



# Effective Health Care Program

---

Comparative Effectiveness Review  
Number 58

## **Noninvasive Technologies for the Diagnosis of Coronary Artery Disease in Women**



Agency for Healthcare Research and Quality  
Advancing Excellence in Health Care • [www.ahrq.gov](http://www.ahrq.gov)

## **Noninvasive Technologies for the Diagnosis of Coronary Artery Disease in Women**

**Prepared for:**

Agency for Healthcare Research and Quality  
U.S. Department of Health and Human Services  
540 Gaither Road  
Rockville, MD 20850  
www.ahrq.gov

**Contract No.** 290-2007-10066-I

**Prepared by:**

Duke Evidence-based Practice Center  
Durham, NC

**Investigators:**

Rowena J. Dolor, M.D., M.H.S., Principal Investigator  
Manesh R. Patel, M.D., Principal Investigator  
Chiara Melloni, M.D., Clinical Investigator  
Ranee Chatterjee, M.D., M.P.H., Clinical Investigator  
Amanda J. McBroom, Ph.D., EPC Project Manager  
Michael D. Musty, EPC Project Coordinator  
Liz Wing, M.A., EPC Editor  
Remy R. Coeytaux, M.D., Ph.D., EPC Investigator  
Adia K. Ross, M.D., M.H.A., Clinical Investigator  
Lori A. Bastian, M.D., M.P.H., Clinical Investigator  
Monique Anderson, M.D., Clinical Investigator  
Andrzej S. Kosinski, Ph.D., Statistical Investigator  
Gillian D. Sanders, Ph.D., EPC Director

This report is based on research conducted by the Duke Evidence-based Practice Center (EPC) under contract to the Agency for Healthcare Research and Quality (AHRQ), Rockville, MD (Contract No. 290-2007-10066-I). The findings and conclusions in this document are those of the authors, who are responsible for its contents; the findings and conclusions do not necessarily represent the views of AHRQ. Therefore, no statement in this report should be construed as an official position of AHRQ or of the U.S. Department of Health and Human Services.

The information in this report is intended to help health care decisionmakers—patients and clinicians, health system leaders, and policymakers, among others—make well-informed decisions and thereby improve the quality of health care services. This report is not intended to be a substitute for the application of clinical judgment. Anyone who makes decisions concerning the provision of clinical care should consider this report in the same way as any medical reference and in conjunction with all other pertinent information, i.e., in the context of available resources and circumstances presented by individual patients.

This report may be used, in whole or in part, as the basis for development of clinical practice guidelines and other quality enhancement tools, or as a basis for reimbursement and coverage policies. AHRQ or U.S. Department of Health and Human Services endorsement of such derivative products may not be stated or implied.

This document is in the public domain and may be used and reprinted without permission except those copyrighted materials that are clearly noted in the document. Further reproduction of those copyrighted materials is prohibited without the specific permission of copyright holders.

Persons using assistive technology may not be able to fully access information in this report. For assistance contact [EffectiveHealthCare@ahrq.hhs.gov](mailto:EffectiveHealthCare@ahrq.hhs.gov).

|  |
|--|
| None of the investigators has any affiliations or financial involvement that conflicts with the material presented in this report. |
|--|

**Suggested Citation:** Dolor RJ, Patel MR, Melloni C, Chatterjee R, McBroom AJ, Musty MD, Wing L, Coeytaux RR, Ross AK, Bastian LA, Anderson M, Kosinski AS, Sanders GD. Noninvasive Technologies for the Diagnosis of Coronary Artery Disease in Women. Comparative Effectiveness Review No. 58. (Prepared by the Duke Evidence-based Practice Center under Contract No. 290-2007-10066-I.) AHRQ Publication No. 12-EHC034-EF. Rockville, MD: Agency for Healthcare Research and Quality. June 2012. [www.effectivehealthcare.ahrq.gov/reports/final.cfm](http://www.effectivehealthcare.ahrq.gov/reports/final.cfm).

## Preface

The Agency for Healthcare Research and Quality (AHRQ) conducts the Effective Health Care Program as part of its mission to organize knowledge and make it available to inform decisions about health care. As part of the Medicare Prescription Drug, Improvement, and Modernization Act of 2003, Congress directed AHRQ to conduct and support research on the comparative outcomes, clinical effectiveness, and appropriateness of pharmaceuticals, devices, and health care services to meet the needs of Medicare, Medicaid, and the Children's Health Insurance Program (CHIP).

AHRQ has an established network of Evidence-based Practice Centers (EPCs) that produce Evidence Reports/Technology Assessments to assist public- and private-sector organizations in their efforts to improve the quality of health care. The EPCs now lend their expertise to the Effective Health Care Program by conducting comparative effectiveness reviews (CERs) of medications, devices, and other relevant interventions, including strategies for how these items and services can best be organized, managed, and delivered.

Systematic reviews are the building blocks underlying evidence-based practice; they focus attention on the strength and limits of evidence from research studies about the effectiveness and safety of a clinical intervention. In the context of developing recommendations for practice, systematic reviews are useful because they define the strengths and limits of the evidence, clarifying whether assertions about the value of the intervention are based on strong evidence from clinical studies. For more information about systematic reviews, see <http://www.effectivehealthcare.ahrq.gov/reference/purpose.cfm>.

AHRQ expects that CERs will be helpful to health plans, providers, purchasers, government programs, and the health care system as a whole. In addition, AHRQ is committed to presenting information in different formats so that consumers who make decisions about their own and their family's health can benefit from the evidence.

Transparency and stakeholder input from are essential to the Effective Health Care Program. Please visit the Web site (<http://www.effectivehealthcare.ahrq.gov>) to see draft research questions and reports or to join an email list to learn about new program products and opportunities for input. Comparative Effectiveness Reviews will be updated regularly.

We welcome comments on this CER. They may be sent by mail to the Task Order Officer named below at: Agency for Healthcare Research and Quality, 540 Gaither Road, Rockville, MD 20850, or by email to [epc@ahrq.hhs.gov](mailto:epc@ahrq.hhs.gov).

Carolyn M. Clancy, M.D.  
Director, Agency for Healthcare Research  
and Quality

Jean Slutsky, P.A., M.S.P.H.  
Director, Center for Outcomes and Evidence  
Agency for Healthcare Research and Quality

Stephanie Chang M.D., M.P.H.  
Director, EPC Program  
Center for Outcomes and Evidence  
Agency for Healthcare Research and Quality

Elisabeth U. Kato, M.D., M.R.P.  
Task Order Officer  
Center for Outcomes and Evidence  
Agency for Healthcare Research and Quality

## Acknowledgments

The authors thank Megan von Isenburg, M.L.S., for help with the literature search and retrieval.

## Key Informants

Bonnie Arkus, R.N.  
Executive Director  
Women's Heart Foundation  
West Trenton, NJ

Javed Butler, M.D., M.P.H.  
Associate Professor of Medicine  
Emory University  
Atlanta, GA

Brenda J. Clark  
Go Red for Women  
American Heart Association

Stacie Daugherty, M.D., M.S.P.H.  
Assistant Professor of Cardiology  
University of Colorado  
Aurora, CO

John F. Heitner, M.D.  
Director of Noninvasive Imaging  
New York Methodist Hospital  
Brooklyn, NY

Mark Hlatky, M.D.  
Director, Stanford-Kaiser Cardiovascular  
Outcomes Research Center  
Stanford University  
Stanford, CA

Neil Jensen, M.B.A., M.H.A.  
Director, Cardiology Networks  
United HealthCare  
Edina, MN

Marcel Salive, M.D., M.P.H.  
Director, Division of Medical and Surgical  
Services  
Centers for Medicare and Medicaid  
Bethesda, MD

## Technical Expert Panel

Matthew J. Budoff, M.D.  
Professor of Medicine  
University of California, Los Angeles  
Los Angeles, CA

Javed Butler, M.D., M.P.H.  
Associate Professor of Medicine  
Emory University  
Atlanta, GA

Stacie Daugherty, M.D., M.S.P.H.  
Assistant Professor of Cardiology  
University of Colorado  
Aurora, CO

John F. Heitner, M.D.  
Director of Noninvasive Imaging  
New York Methodist Hospital  
Brooklyn, NY

Mark Hlatky, M.D.  
Director, Stanford-Kaiser Cardiovascular  
Outcomes Research Center  
Stanford University  
Stanford, CA

Neil Jensen, M.B.A., M.H.A.  
Director, Cardiology Networks  
United HealthCare  
Edina, MN

Marcel Salive, M.D., M.P.H.  
Director, Division of Medical and Surgical  
Services  
Centers for Medicare and Medicaid  
Bethesda, MD

Subha V. Raman, M.D., M.S.  
Cardiovascular Imaging Research Center  
Ohio State University Medical Center  
Columbus, OH

Malissa J. Wood, M.D.  
Assistant Professor Medicine  
Harvard Medical School  
Boston, MA

## **Peer Reviewers**

Elizabeth Barrett-Connor, M.D.  
Professor and Division Chief of  
Epidemiology  
University of California, San Diego  
San Diego, CA

Kavitha M. Chinnaiyan, M.D.  
Medical Director of Noninvasive Cardiology  
Education  
Beaumont Heart Center  
Royal Oak, MI

Nakela L. Cook, M.D., M.P.H.  
Division of Cardiology  
Washington Hospital Center  
Washington, DC

Jean McSweeney, Ph.D., R.N.  
Associate Dean for Research  
University of Arkansas for Medical Sciences  
Little Rock, AR

Lori Mosca, M.D., Ph.D., M.P.H.  
Professor of Medicine  
Director of Preventive Cardiology  
Columbia University Medical Center  
New York, NY

Frank John Rybicki III, M.D., Ph.D.  
Associate Professor of Radiology  
Brigham and Women's Hospital  
Boston, MA

# Noninvasive Technologies for the Diagnosis of Coronary Artery Disease in Women

## Structured Abstract

**Objectives:** To conduct a systematic review of the medical literature assessing (1) accuracy of noninvasive technologies (NITs) for diagnosing coronary artery disease (CAD) in women with symptoms suspicious for CAD, (2) predictors affecting test accuracy, (3) ability of NITs to provide risk stratification, prognostic information, inform decisionmaking about treatment options, and affect clinical outcomes, and (4) risks to women undergoing these tests.

**Data Sources:** MEDLINE<sup>®</sup>, PubMed<sup>®</sup>, Embase<sup>®</sup>, and Cochrane Database of Systematic Reviews.

**Review Methods:** Studies published in English through September 2011 with sex-specific outcomes comparing exercise/stress electrocardiography (ECG), echocardiography (ECHO), single proton emission computed tomography (SPECT), cardiac magnetic resonance (CMR), or coronary computed tomography angiography (coronary CTA) with another NIT, or with coronary angiography. We ran separate meta-analyses of the accuracy of each NIT modality compared with coronary angiography on the no known and mixed CAD populations in women and in men.

**Results:** A total of 104 comparative studies (110 articles) were included. For women with no known CAD, the summary of accuracy for each NIT modality compared with coronary angiography was ECG (29 studies), sensitivity 62 percent, specificity 68 percent; ECHO (14 studies), sensitivity 79 percent, specificity 83 percent; SPECT (14 studies), sensitivity 81 percent, specificity 78 percent; CMR (5 studies), sensitivity 72 percent, specificity 84 percent; and CTA (5 studies), sensitivity 94 percent, specificity 87 percent. Compared with men, in women ECG and coronary CTA modalities were both less sensitive and less specific. The ECHO and SPECT modalities, although less sensitive, appeared to be more specific in women. The lower specificity of the ECG modality in women was the only statistically significant difference. Strength of evidence was high for ECG, ECHO, and SPECT and low for CMR and coronary CTA compared with coronary angiography in women. Eleven comparative studies examined predictors of diagnostic accuracy in women such as postmenopausal status, race/ethnicity, heart size, beta blocker use, and pretest probability; insufficient evidence was available to draw conclusions about predictors that affect accuracy. Eight studies assessed risk stratification and prognostic factors, two studies assessed treatment decisionmaking, and four studies provided comparative clinical outcomes. There is insufficient evidence on the comparative effectiveness of NITs to provide risk stratification, prognostic information, treatment decisionmaking, or impact clinical outcomes in women. Thirteen comparative studies reported risks. Of these, four studies of coronary CTA showed a higher mean effective radiation dose and attributable risk of cancer incidence in women compared with men; however, radiation safety issues were not discussed in other NIT modalities with radiation exposure. Thus, there was insufficient evidence regarding the comparative risks of various NIT modalities in women.

**Conclusions:** This systematic review provides evidence for the summary sensitivities and specificities of exercise/stress ECG, ECHO, SPECT, CMR, and coronary CTA compared with coronary angiography used for diagnosing CAD in women. There was limited or insufficient evidence from comparative studies to define the influence of clinical and demographic factors on NIT diagnostic accuracy, risk stratification, prognostic information, treatment decisions, clinical outcomes, and harms in women.



# Contents

|  |      |
|--|------|
| <b>Executive Summary</b> .....   | ES-1 |
| <b>Introduction</b> .....  | 1    |
| Background .....   | 1    |
| Types of Noninvasive Technologies .....  | 2    |
| Uncertainties Surrounding Noninvasive Diagnosis of CAD in Women .....                                      | 5    |
| Relevance .....  | 6    |
| Scope and Key Questions .....  | 6    |
| <b>Methods</b> .....   | 8    |
| Topic Development and Refinement .....   | 8    |
| Analytic Framework .....   | 8    |
| Literature Search Strategy .....   | 9    |
| Sources Searched .....   | 9    |
| Process for Study Selection .....  | 10   |
| Screening for Inclusion and Exclusion .....  | 10   |
| Data Extraction and Data Management .....  | 12   |
| Individual Study Quality Assessment .....  | 13   |
| Data Synthesis .....   | 14   |
| Grading the Body of Evidence .....   | 16   |
| Peer Review and Public Commentary .....  | 16   |
| <b>Results</b> .....   | 17   |
| Key Question 1: Diagnostic Accuracy of NITs .....  | 20   |
| Key Points .....   | 20   |
| Detailed Synthesis .....   | 21   |
| KQ 1 Summary .....   | 69   |
| Key Question 2: Predictors of Diagnostic Accuracy .....  | 73   |
| Key Points .....   | 73   |
| Detailed Synthesis .....   | 73   |
| KQ 2 Summary .....   | 87   |
| Key Question 3: Use of NITs To Improve Risk Stratification, Decisionmaking, and<br>Clinical Outcomes ..... | 87   |
| Key Points .....   | 87   |
| Detailed Synthesis .....   | 87   |
| KQ 3 Summary .....   | 98   |
| Key Question 4: Safety Concerns and Risks .....  | 98   |
| Key Points .....   | 98   |
| Detailed Synthesis .....   | 98   |
| KQ 4 Summary .....   | 110  |
| Summary and Discussion .....   | 110  |
| KQ 1: Diagnostic Accuracy of NITs .....  | 110  |

|  |            |
|--|------------|
| KQ 2: Predictors of Diagnostic Accuracy .....  | 115        |
| KQ 3: Improving Risk Stratification, Decisionmaking, and Outcomes .....  | 115        |
| KQ 4: Safety Concerns .....  | 114        |
| Discussion .....   | 116        |
| Limitations of This Review .....   | 118        |
| Conclusions .....  | 119        |
| <b>Future Research</b> .....   | <b>123</b> |
| <b>References</b> .....  | <b>125</b> |
| <b>Abbreviations</b> .....   | <b>133</b> |
| <b>Tables</b>  |            |
| Table A. ....  | ES-6       |
| Table B. ....  | ES-12      |
| Table 1. Summary of Inclusion and Exclusion Criteria .....   | 11         |
| Table 2. Summary of Accuracy Data Evaluating ECG for Diagnosing CAD .....  | 31         |
| Table 3. Summary of Accuracy Data Evaluating ECHO for Diagnosing CAD .....   | 43         |
| Table 4. Summary of Accuracy Data Evaluating SPECT for Diagnosing CAD .....  | 52         |
| Table 5. Summary of Accuracy Data Evaluating CMR for Diagnosing CAD .....  | 60         |
| Table 6. Summary of Accuracy Data Evaluating Coronary CTA for Diagnosing CAD .....   | 67         |
| Table 7. Summary of Accuracy of Nits Compared With Coronary Angiography for<br>Diagnosing CAD in Women .....                                 | 70         |
| Table 8. Summary of Accuracy of Nits for Diagnosing CAD in Men Compared With<br>Women from Mixed Populations .....                           | 72         |
| Table 9. Age and Menopausal Status as a Predictor .....  | 75         |
| Table 10. Race/Ethnicity as a Predictor .....  | 75         |
| Table 11. Heart Size as a Predictor .....  | 76         |
| Table 12. Other Potential Predictors .....   | 78         |
| Table 13. Summary Table for ECG.....   | 83         |
| Table 14. Summary Table for ECHO.....  | 84         |
| Table 15. Summary Table for SPECT.....   | 85         |
| Table 16. Summary Table for CMR.....   | 85         |
| Table 17. Summary of Findings for KQ 3.....  | 91         |
| Table 18. Adverse Effects of Different Nits for Screening of CAD in Women .....  | 103        |
| Table 19. Summary of Accuracy of Nits for Diagnosing CAD in Women .....  | 111        |
| Table 20. GRADE Table for Accuracy of NIT Modalities in Women With no Known<br>CAD.....  | 114        |
| Table 21. Summary of Key Findings .....  | 116        |
| <b>Figures</b>   |            |
| Figure A. Analytic Framework .....   | ES-4       |
| Figure B. Literature Flow Diagram.....   | ES-11      |
| Figure C. Summary of Accuracy of NITs Compared With Coronary Angiography for<br>Diagnosing CAD in Women With no Known CAD (All Studies)..... | ES-13      |

|   |    |
|---|----|
| Figure 1. Analytic Framework .....  | 9  |
| Figure 2. Literature Flow Diagram.....  | 18 |
| Figure 3. QUADAS Elements Used To Rate Diagnostic Accuracy .....                                      | 19 |
| Figure 4. Accuracy of ECG in Women With no Known CAD.....   | 23 |
| Figure 5. SROC Curve for ECG in Women With no Known CAD.....  | 25 |
| Figure 6. Accuracy of ECG in 10 Good-Quality Studies in Women With no Known CAD...                    | 25 |
| Figure 7. SROC Curve for ECG in 10 Good-Quality Studies in Women With no Known<br>CAD.....            | 26 |
| Figure 8. Accuracy of ECG in Women from Mixed Populations .....                                       | 27 |
| Figure 9. SROC Curve for ECG in Women from Mixed Populations.....                                     | 28 |
| Figure 10. Accuracy of ECG in 13 Good-Quality Studies in Women from Mixed<br>Populations .....        | 29 |
| Figure 11. SROC Curve for ECG in 13 Good-Quality Studies in Women from Mixed<br>Populations .....     | 30 |
| Figure 12. Accuracy of ECHO in Women With no Known CAD.....   | 36 |
| Figure 13. SROC Curve for ECHO in Women With no Known CAD.....  | 37 |
| Figure 14. Accuracy of ECHO in Five Good-Quality Studies in Women With no Known<br>CAD.....           | 38 |
| Figure 15. SROC Curve for ECHO in Five Good-Quality Studies in Women With no<br>Known CAD.....        | 38 |
| Figure 16. Accuracy of ECHO in Women from Mixed Populations .....                                     | 39 |
| Figure 17. SROC Curve for ECHO in Women from Mixed Populations.....                                   | 40 |
| Figure 18. Accuracy of ECHO in Eight Good-Quality Studies in Women from Mixed<br>Populations .....    | 41 |
| Figure 19. SROC Curve for ECHO in Eight Good-Quality Studies in Women from Mixed<br>Populations ..... | 41 |
| Figure 20. Accuracy of SPECT in Women With no Known CAD.....  | 45 |
| Figure 21. SROC Curve for SPECT in Women With no Known CAD.....                                       | 46 |
| Figure 22. Accuracy of SPECT in Four Good-Quality Studies in Women With no Known<br>CAD.....          | 47 |
| Figure 23. SROC Curve for SPECT in Four Good-Quality Studies in Women With no<br>Known CAD.....       | 47 |
| Figure 24. Accuracy of SPECT in Women from Mixed Populations .....                                    | 49 |
| Figure 25. SROC Curve for SPECT in Women from Mixed Populations.....                                  | 50 |
| Figure 26. Accuracy of SPECT in 10 Good-Quality Studies in Women from Mixed<br>Populations .....      | 51 |
| Figure 27. SROC Curve for SPECT in 10 Good-Quality Studies in Women from Mixed<br>Populations .....   | 51 |
| Figure 28. Accuracy of CMR in Women With no Known CAD .....   | 55 |
| Figure 29. SROC Curve for CMR in Women With no Known CAD.....   | 56 |
| Figure 30. Accuracy of CMR in Women from Mixed Populations .....                                      | 57 |
| Figure 31. SROC Curve for CMR in Women from Mixed Populations.....                                    | 57 |
| Figure 32. Accuracy of CMR in Five Good-Quality Studies in Women from Mixed<br>Populations .....      | 58 |
| Figure 33. SROC Curve for CMR in Five Good-Quality Studies in Women from Mixed<br>Populations .....   | 58 |

|   |     |
|---|-----|
| Figure 34. Accuracy of Coronary CTA in Women With no Known CAD.....   | 61  |
| Figure 35. SROC Curve for Coronary CTA in Women With no Known CAD.....  | 62  |
| Figure 36. Accuracy of Coronary CTA in Women from Mixed Populations.....  | 63  |
| Figure 37. SROC Curve for Coronary CTA in Women from Mixed Populations.....   | 64  |
| Figure 38. Accuracy of Coronary CTA in Four Good-Quality Studies in Women from<br>Mixed Populations.....                                      | 65  |
| Figure 39. SROC Curve for Coronary CTA in Four Good-Quality Studies in Women from<br>Mixed Populations.....                                   | 65  |
| Figure 40. Summary of Accuracy of Nits Compared With Coronary Angiography for<br>Diagnosing CAD in Women With no Known CAD (All Studies)..... | 71  |
| Figure 41. Summary of Accuracy of Nits for Diagnosing CAD in Women With no Known<br>CAD (All Studies).....                                    | 112 |

## **Appendixes**

- Appendix A. Exact Search Strings
- Appendix B. Data Abstraction Elements
- Appendix C. List of Included Studies
- Appendix D. Quality and Applicability of Included Studies
- Appendix E. List of Excluded Studies

# Executive Summary

## Background

Cardiovascular disease is the leading cause of mortality for women in the United States.<sup>1</sup> Coronary heart disease—which includes coronary artery (or atherosclerotic) disease (CAD), myocardial infarction (MI), acute coronary syndromes, and angina—is the largest subset of this mortality.<sup>1</sup> According to the American Heart Association (AHA), approximately one in three female adults has some form of cardiovascular disease. Since 1984, the number of deaths attributed to cardiovascular disease in women has exceeded that in men, reaching 454,613 in 2005—more than deaths from all forms of cancer combined.<sup>2</sup> It is estimated that 8.1 million women alive today have a history of heart attack, angina pectoris (chest pain or discomfort caused by reduced blood supply to the heart muscle), or both, and experts predict that in 2010 alone an estimated 370,000 women will have a new or recurrent MI. Overall, women who have had an acute MI—particularly those older than 55 years of age—have a worse prognosis than men, with a greater recurrence of MI and higher mortality.<sup>1</sup> More women (5.5 million) than men (4.3 million) have angina in total numbers. However, the prevalence of CAD in women with chest pain is about 50 percent, compared with 80 percent in men, which complicates diagnosis in women.<sup>3</sup>

The AHA suggests there is evidence showing that women at risk for CAD are less often referred for the appropriate diagnostic test than are men.<sup>1</sup> Coronary anatomy and pathology have traditionally been defined and identified by invasive, catheter-based x-ray angiography, also referred to as coronary angiography.<sup>4</sup> In this invasive procedure, a catheter is inserted into the femoral, brachial, or radial artery and passed up through the aorta to directly engage the right and left coronary arteries; an iodinated contrast agent is then injected into each artery while digital x-ray images are recorded.<sup>4</sup> The major benefits of invasive coronary angiography over noninvasive techniques are that the use of a catheter makes it possible to see the coronary arteries with greater anatomic precision and resolution and to combine diagnosis and treatment in a single procedure. The limitations of the procedure include the invasive nature of the test and the limited data on the functional impact of a luminal obstruction. These limitations are generally considered to be minor when compared with the benefits of the procedure, and coronary angiography is now the reference (gold) standard for clinical care of patients who have chest pain suggestive of CAD.

Coronary angiography, however, is not risk-free.<sup>4</sup> Arterial bleeding can occur at the access site, and manipulation of the catheter within the aorta and coronary arteries may cause an atherosclerotic embolus that, in turn, could result in stroke or heart attack. Separation of material from the inner lining of the artery may also cause a blockage downstream of the catheter tip. The contrast agent used during the procedure to visualize the coronary arteries may cause anaphylaxis, renal impairment, or injury, and there is radiation exposure during the digital x-ray imaging. Although it is a rare occurrence, the catheter can puncture an artery and cause internal bleeding.

Coronary angiography is generally indicated in patients who have chest pain and are at high risk for CAD. For intermediate-risk patients, clinicians have a wide range of noninvasive diagnostic modalities to choose from, with wide variability in reported sensitivities and specificities. Noninvasive technologies (NITs) are especially important options for patients who have contraindications to invasive catheterization, or for those who would be put at higher risk for complications with invasive screening.<sup>4</sup>

## Types of Noninvasive Technologies

NITs can assess functional status (i.e., ischemia or no ischemia) or visualize anatomic abnormalities (i.e., no CAD, nonobstructive CAD, or obstructive CAD). Types of NITs include the following:

- Functional modalities:
  - Exercise/stress electrocardiography (ECG) exercise/stress or resting
  - Exercise/stress echocardiography (ECHO) with or without a contrast agent
  - Exercise/stress radionuclide myocardial perfusion imaging, including single proton emission computed tomography (SPECT) and positron emission tomography (PET)
- Anatomic modalities:
  - Stress myocardial perfusion and wall motion magnetic resonance imaging (CMR)
  - Coronary computed tomography angiography (coronary CTA)

The AHA and the American College of Cardiology (ACC) recommend that women with suspected CAD should be classified as either symptomatic or asymptomatic and further classified as being at low, intermediate, or high risk for the disease to guide the decision about which diagnostic test to use first.<sup>1</sup> In 2005, the AHA developed a consensus statement on the role of noninvasive testing in the clinical evaluation of women with suspected CAD. In this statement, the AHA recommended that women who are symptomatic and at intermediate to high risk of having CAD should undergo noninvasive diagnostic studies (i.e., exercise electrocardiography and cardiac imaging studies) and that those who are asymptomatic and at low risk of CAD should not undergo cardiac imaging studies.<sup>1</sup> The AHA consensus statement was a thorough synopsis of the extant literature regarding the diagnosis of CAD in women with expert-guided recommendations for the workup of symptomatic women but did not include a comparative effectiveness review of the accuracy of the various NIT modalities in women.

## Objectives

The goal of this comparative effectiveness report was to conduct a systematic review of the peer-reviewed medical literature assessing (1) the accuracy of different NITs for diagnosing CAD in women with symptoms suspicious of CAD, (2) the predictors affecting test accuracy, (3) the ability to provide risk stratification and prognostic information, inform decisionmaking about treatment options, and affect clinical outcomes, and (4) the safety concerns and risks to women undergoing these tests. The following Key Questions (KQs) were considered in this comparative effectiveness review:

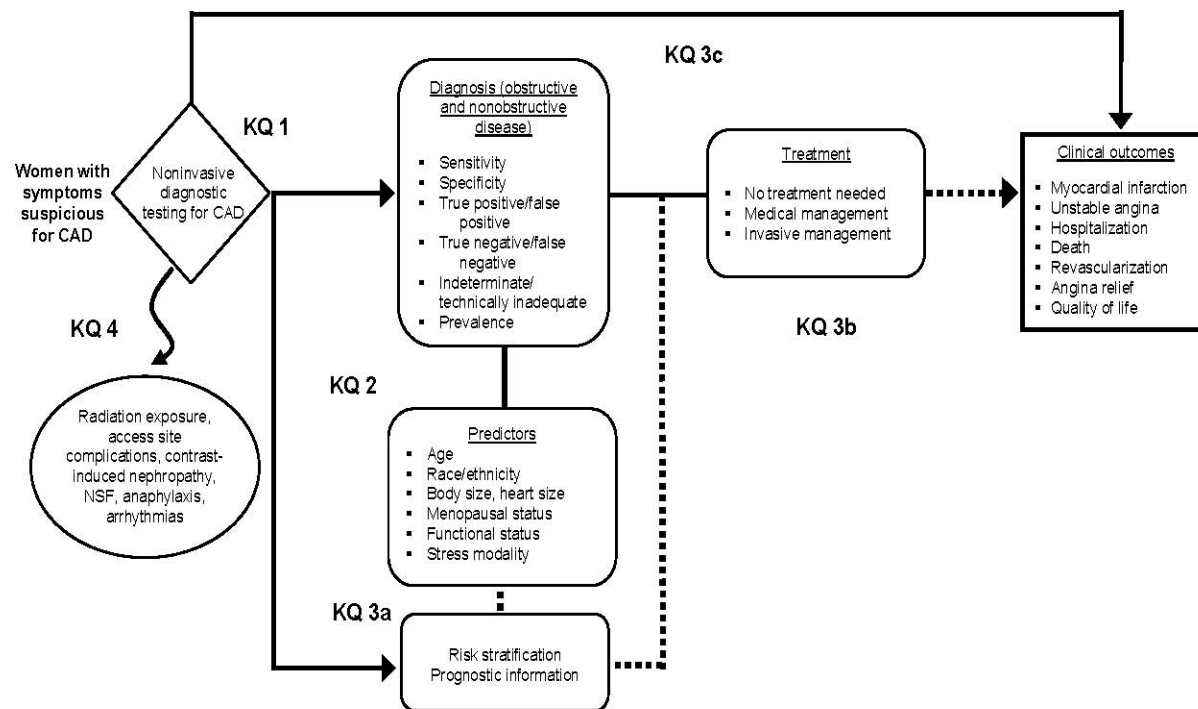
- **KQ 1.** What is the accuracy of one NIT in diagnosing obstructive and nonobstructive CAD when compared with another NIT or with coronary angiography in women with symptoms suspicious for CAD?
  - Exercise ECG stress test, including resting ECG technology (e.g., multifunctional cardiogram)

- Exercise/stress ECHO with or without a contrast agent
- Exercise/stress radionuclide myocardial perfusion imaging, including SPECT and PET
- CMR imaging
- Coronary CTA
- **KQ 2.** What are the predictors of diagnostic accuracy (e.g., age, race/ethnicity, body size, heart size, menopausal status, functional status, stress modality) of different NITs in women?
- **KQ 3.** Is there evidence that the use of NITs (when compared with other NITs or with coronary angiography) in women improves:
  - **KQ 3a.** Risk stratification/prognostic information?
  - **KQ 3b.** Decisionmaking regarding treatment options (e.g., revascularization, optimal medical therapy)?
  - **KQ 3c.** Clinical outcomes (e.g., death, myocardial infarction, unstable angina, hospitalization, revascularization, angina relief, quality of life)?
- **KQ 4.** Are there significant safety concerns/risks (i.e., radiation exposure, access site complications, contrast agent-induced nephropathy, nephrogenic systemic fibrosis, anaphylaxis, arrhythmias) associated with the use of different NITs to diagnose CAD in women with symptoms suspicious for CAD?

## Analytic Framework

Figure A shows the analytic framework for the systematic review of NITs for the diagnosis of CAD in women.

Figure A. Analytic framework



Abbreviations: CAD = coronary artery disease; KQ = Key Question; NSF = nephrogenic systemic fibrosis

## Methods

### Input From Stakeholders

The Evidence-based Practice Center (EPC) followed AHRQ's recommended methodology, described in Methods Guide for Effectiveness and Comparative Effectiveness Reviews,<sup>5</sup> for literature search strategies, inclusion/exclusion of studies, abstract screening, data abstraction and management, assessment of methodological quality of individual studies, data synthesis, and grading of evidence for each KQ.

During the topic refinement stage, we solicited input from Key Informants, representing clinicians (cardiology, primary care, cardiac imaging), patients, scientific experts, and Federal agencies to help define the KQs. The KQs were then posted for public comment for 30 days, and the comments received were considered in the development of the research protocol. We next convened a Technical Expert Panel (TEP), comprising clinical, content, and methodological experts, to provide input in defining populations, interventions, comparisons, or outcomes as well as identifying particular studies or databases to search. The Key Informants and members of



the TEP were required to disclose any financial conflicts of interest greater than \$10,000 and any other relevant business or professional conflicts of interest. Any potential conflicts of interest were balanced or mitigated. Neither Key Informants nor members of the TEP did analysis of any kind and did not contribute to the writing of the report.

## **Data Sources and Selection**

We included studies published in English from January 1, 2000, through September 12, 2011. Search strategies were specific to each database in order to retrieve the articles most relevant to the KQs. Our search strategy used the National Library of Medicine's medical subject headings (MeSH) keyword nomenclature developed for MEDLINE<sup>®</sup> and adapted for use in other databases. We used PubMed<sup>®</sup>, Embase<sup>®</sup>, the Cochrane Database of Systematic Reviews, and the Cochrane Central Registry of Controlled Trials for our literature search. We also searched the grey literature of study registries and conference abstracts for relevant articles from completed trials, including Clinicaltrials.gov; metaRegister of Controlled Trials; ClinicalStudyResults.org; WHO: International Clinical Trials Registry Platform Search Portal; CSA Conference Papers Index; and Scopus. The exact search strings used in our strategy are given in Appendix A of the full report. Reference lists of articles applicable to the relevant KQs of previous AHRQ reports on this topic<sup>6,7</sup> and from identified systematic reviews and meta-analyses were manually hand-searched and cross-referenced against our library, and additional manuscripts were retrieved. All citations were imported into an electronic bibliographic database (EndNote<sup>®</sup> Version X4; Thomson Reuters, Philadelphia, PA).

We developed a list of article inclusion and exclusion criteria for the KQs (Table A). Using the prespecified inclusion and exclusion criteria, titles and abstracts were examined independently by two reviewers for potential relevance to the KQs. Articles included by any reviewer underwent full-text screening. At the full-text screening stage, two independent reviewers read each article to determine if it met eligibility criteria. At the full-text review stage, paired researchers independently reviewed the articles and indicated a decision to "include" or "exclude" the article for data abstraction. When the paired reviewers arrived at different decisions about whether to include or exclude an article, we reconciled the difference through a third-party arbitrator. Articles meeting our eligibility criteria were included for data abstraction. Relevant systematic review articles, meta-analyses, and methods articles were flagged for hand-searching and cross-referencing against the library of citations identified through electronic database searching.

**Table A. Summary of inclusion and exclusion criteria**

| Study Characteristic | Inclusion Criteria   | Exclusion Criteria  |
|----------------------|--|---|
| Study design         | <p>An article was included if the following two criteria were met:</p> <p>Original data or related methodology paper of an included article</p> <p>Randomized controlled trial, prospective or retrospective observational study, or registry</p>  | <p>Editorial</p> <p>Not a systematic review</p> <p>Letter to editor</p> <p>Case series</p> <p>Review article, meta-analysis, or methods paper of an excluded article</p> <p>Not peer reviewed</p>   |
| Population           | <p>Study included adult women (age <math>\geq</math> 18 years of age) who present symptoms of symptoms suspicious for CAD (e.g., exertional dyspnea, shortness of breath, and/or angina) with or without a known diagnosis of CAD; data for women must be presented separately from data for men</p>   | <p>All subjects were &lt; 18 years of age, or some subjects were under &lt; 18 but results were not broken down by age</p> <p>No patients had symptomatic chest pain (i.e., an asymptomatic population), or some of the patients had symptomatic chest pain but results were not reported separately for this subgroup</p> <p>All patients were known to have CAD and were not being tested for chest pain symptoms (e.g., postrevascularization testing to assess for persistent ischemia)</p> |
| Interventions        | <p>NITs for the diagnosis of obstructive and nonobstructive CAD included:</p> <p>Exercise electrocardiogram stress test</p> <p>Resting electrocardiogram technology</p> <p>Exercise/stress echocardiography with or without a contrast agent</p> <p>Exercise/stress radionuclide myocardial perfusion imaging, including single proton emission computed tomography and positron emission tomography</p> <p>Stress myocardial perfusion and wall motion magnetic resonance imaging</p> <p>Coronary computed tomography angiography</p> | <p>Coronary artery calcium scoring by electron beam computed tomography since this modality is often used to screen asymptomatic patients for CAD</p>   |

**Table A. Summary of inclusion and exclusion criteria (continued)**

| Study Characteristic  | Inclusion Criteria   | Exclusion Criteria   |
|-----------------------|--|--|
| Comparators           | Another NIT or coronary angiography  | Study did not compare one NIT with another, or with coronary angiography   |
| Outcomes              | Primary outcome—accurate diagnosis of obstructive and nonobstructive CAD   | Outcomes not related to diagnostic accuracy for detecting CAD  |
|                       | KQ 1 patient-level outcomes:<br>Sensitivity<br>Specificity<br>True positive, false negative, true negative, false positive<br>Indeterminate or technically inadequate results<br>Prevalence  | Vessel-based outcomes<br>Outcomes of women not reported separately from total population   |
|                       | KQ 2 outcomes: Predictors of diagnostic accuracy—age, race/ethnicity, body size, heart size, menopausal status, functional status, stress modality   | Outcomes of women not reported separately from total population  |
|                       | KQ 3 outcomes:<br>Risk stratification/prognostic information<br>Treatment —none, medical therapy, percutaneous coronary intervention, or coronary artery bypass surgery<br>Clinical outcomes—death, myocardial infarction, unstable angina, hospitalization, revascularization, angina relief, quality of life | Outcomes of women not reported separately from total population  |
|                       | KQ 4 outcomes: Safety and adverse events—radiation exposure, access site complications, contrast agent-induced nephropathy, nephrogenic systemic fibrosis, anaphylaxis, arrhythmias—and how these events varied by demographic factors   | Outcomes of women not reported separately from total population  |
| Setting               | Inpatient or outpatient settings, primarily primary care and cardiology clinics  | None   |
| Publication languages | English only   | Given the high volume of English-language publications (including the majority of known important studies), non-English articles were excluded |

Abbreviations: CAD = coronary artery disease; KQ = Key Question; NIT = noninvasive technology

## Data Extraction and Quality Assessment

The investigative team created forms for abstracting the data elements for the KQs. Based on their clinical and methodological expertise, two researchers were assigned to abstract data from the eligible articles pertaining to the research questions. One researcher abstracted the data, and the second overread the article and the accompanying abstraction form to check for accuracy and completeness. Disagreements were resolved by consensus or by obtaining a third reviewer's opinion if consensus was not reached by the first two researchers. To aid in both reproducibility and standardization of data collection, researchers received data abstraction instructions directly on each form created specifically for this project with the DistillerSR data synthesis software program (Evidence Partners Inc., Manotick, ON, Canada). We designed these forms to collect the data required to evaluate the specified eligibility criteria for inclusion in this review as well as to collect demographics and data needed to determine outcomes (intermediate outcomes, health outcomes, and safety outcomes). Appendix B of the full report lists the elements used in the data abstraction forms.

Appendix C of the full report contains a bibliography of all studies included in this review, organized alphabetically by author. When appropriate, methods articles providing additional detail were considered when abstracting data for an included study.

The studies included in this comparative effectiveness review were assessed on the basis of the quality of their reporting of relevant data. We evaluated the quality of individual studies using the approach described in AHRQ's Methods Guide for Effectiveness and Comparative Effectiveness Reviews (hereafter referred to as the Methods Guide).<sup>5</sup> To assess study quality, we (1) classified the study design, (2) applied predefined criteria for quality and critical appraisal, and (3) made a summary judgment of the study's quality. To evaluate methodological quality, we applied criteria for each study type that were derived from the core elements described in the Methods Guide<sup>5</sup> and from QUADAS,<sup>8</sup> a tool for the quality assessment of studies of diagnostic accuracy included in systematic reviews. To indicate the summary judgment of the quality of the individual studies, we used the summary ratings of Good, Fair, and Poor based on the study's adherence to well-accepted standard methodologies (such as QUADAS) and adequate reporting standards.

We used data abstracted on the population studied, the intervention and comparator, the outcomes measured, settings, and timing of assessments to identify specific issues that may have limited the applicability of individual studies or a body of evidence as recommended in the Methods Guide.<sup>5,9</sup> We used these data to evaluate the applicability to clinical practice, paying special attention to study eligibility criteria, demographic features of the enrolled population in comparison with the target population, the intervention used in comparison with technologies currently in use, and clinical relevance and timing of the outcome measures. We summarized issues of applicability qualitatively. Appendix D of the full report summarizes our assessment of the quality and applicability for each included study as well as the assessed QUADAS quality scores for diagnostic accuracy.

## Data Synthesis and Analysis

We summarized the primary literature by abstracting relevant continuous data (e.g., age, sensitivity, specificity, event rates) and categorical data (e.g., race/ethnicity, presence of CAD). Data for patients with no known diagnosis of CAD were collected and analyzed separately from data for mixed CAD populations that included patients with and without known CAD. We then

determined the feasibility of completing a quantitative synthesis (i.e., summary receiver operating characteristic [SROC] curves for diagnostic accuracy or meta-analysis for other outcomes). The feasibility of a meta-analysis or SROC curve depended on the volume of relevant literature, the homogeneity of the studies in terms of the populations studied, the interventions included or the outcomes assessed, and the completeness of the results reporting. For each SROC calculation, we ran separate analyses of the accuracy of each NIT modality compared with coronary angiography on the no-known CAD and mixed CAD populations using random-effects models to quantitatively synthesize the available evidence. In our primary analyses, we evaluated these performance characteristics in the population of women who had no previously known CAD. In secondary analyses, we explored a broader patient population by including those studies that had women from mixed populations of known and no known CAD. We also assessed the impact on our findings if, in each population, we restricted our analyses to those studies that were assessed to be good quality. We then compared the performance characteristics of the NIT modalities with each other in a generalized linear mixed model. In a final exploratory analysis, we evaluated the test performance of the modalities in women compared with men in a similar generalized linear model with sex as a covariate. We presented summary estimates and confidence intervals (CIs).

For synthesizing the accuracy data for studies included in our assessment of KQ 1, we used the following approach as advocated by Leeflang, et al.<sup>10</sup> This approach allows the paired nature of sensitivity and specificity and randomness between studies to be taken into account. The analyses are based on true positive (TP), false negative (FN), false positive (FP), and true negative (TN) frequencies abstracted from relevant publications. Estimated study specific sensitivity ( $TP/[TP+FN]$ ) and specificity ( $TN/[TN+FP]$ ) values are displayed in paired forest plots together with exact 95% CIs.<sup>11</sup> The fixed-effects estimates and their variance–covariance matrix provided (after reverse logit transformation) summary sensitivity and specificity values and a joint confidence region (dotted oval shape on figures) as well as separate CIs for summary sensitivity and specificity as presented on figures and forest plots in the report. We used the Rutter and Gatsonis<sup>12</sup> SROC curve as described by Arends, et al.,<sup>13</sup> and it is presented in figures as a solid line over the range of the available data.

## Grading the Body of Evidence

The strength of evidence for each Key Question was assessed using the approach described in AHRQ’s Methods Guide on Medical Test Reviews for grading the evidence related to the diagnostic accuracy of the NITs (KQ 1),<sup>14</sup> and the Methods Guide for Effectiveness and Comparative Effectiveness Reviews for grading the evidence related to the other Key Questions (KQs 2–4).<sup>5,15</sup> The evidence was evaluated using the four required domains: risk of bias (low, medium, or high), consistency (consistent, inconsistent, or unknown/not applicable), directness (direct or indirect), and precision (precise or imprecise). Additionally, when appropriate, the studies were evaluated for the presence of confounders that would diminish an observed effect, the strength of association (magnitude of effect), and publication bias. The strength of evidence was assigned an overall grade of High, Moderate, Low, or Insufficient according to the following four-level scale:

- **High**—High confidence that the evidence reflects the true effect. Further research is very unlikely to change our confidence in the estimate of effect.

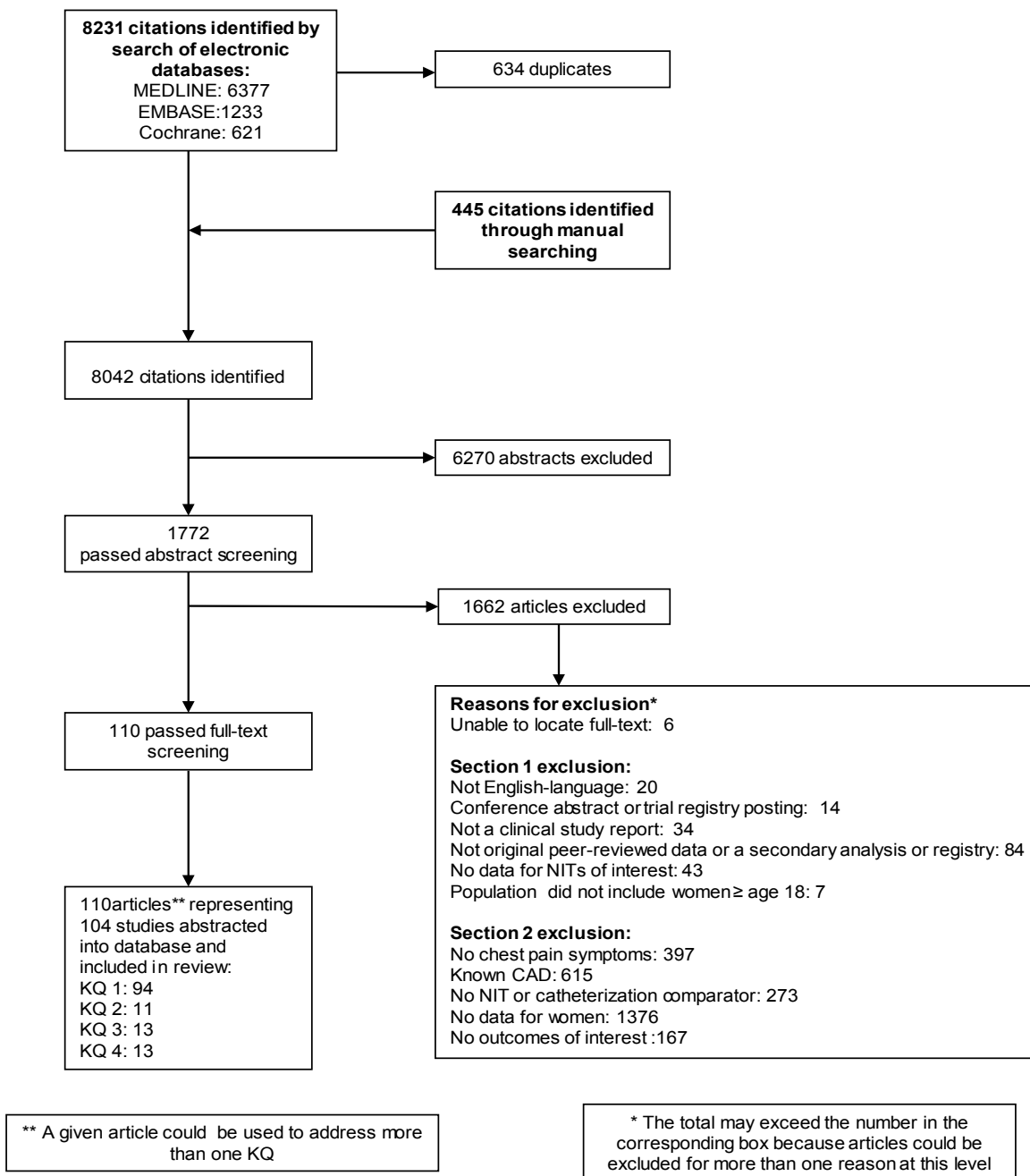
- Moderate—Moderate confidence that the evidence reflects the true effect. Further research may change our confidence in the estimate of effect and may change the estimate.
- Low—Low confidence that the evidence reflects the true effect. Further research is likely to change the confidence in the estimate of effect and is likely to change the estimate.
- Insufficient—Evidence either is unavailable or does not permit estimation of effect.

## Results

The flow of articles through the literature search and screening process is depicted in Figure B. Of the 8,231 citations identified by our searches, 634 were duplicates. A manual search identified an additional 445 citations for a total of 8,042 citations. After applying inclusion/exclusion criteria at the title-and-abstract level, 1,772 full-text articles were retrieved and screened. Of these, 1,662 articles were excluded at the full-text screening stage. Of these, we excluded 1,376 (83 percent) for not reporting data on women and 615 (37 percent) for looking only at a population with known CAD. (Note that an article may have been excluded for more than one reason.) The final set comprised 110 articles representing 104 studies.

Of the 104 studies, 1 was an RCT, 79 were prospective observational, and 24 were retrospective observational with study cohorts comprising individuals who presented for NIT testing and received diagnostic coronary angiography (100 studies) or another NIT modality only (4 studies). The four studies without coronary angiography compared ECHO with ECG<sup>16,17</sup> or ECG with SPECT.<sup>18,19</sup> Three of these studies were applicable to KQ 3,<sup>16-18</sup> and one was applicable to KQ 2.<sup>19</sup> Of the 94 studies included in the KQ 1 results, 5 reported NIT versus NIT comparisons in addition to coronary angiography.<sup>20-24</sup>

**Figure B. Literature flow diagram**



Abbreviations: CAD = coronary artery disease; KQ = Key Question; NIT = noninvasive technology; SR = systematic review

## Summary of Key Findings

We analyzed the results by study population (no known CAD and mixed CAD populations) and by study quality (good quality rating). Table B and Figure C show the summary sensitivities and specificities for each NIT modality. Table C summarizes our key findings.

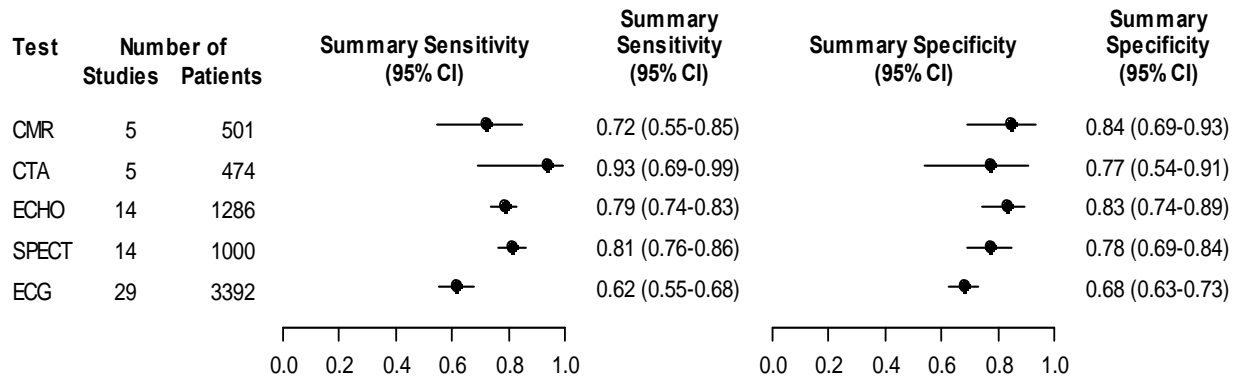
**Table B. Summary of accuracy of NITs compared with coronary angiography for diagnosing CAD in women**

| Modality     | Population       | Quality of Included Studies | Number of Studies | Number of Women | Summary Sensitivity (95% CI) | Summary Specificity (95% CI) |
|--------------|------------------|-----------------------------|-------------------|-----------------|------------------------------|------------------------------|
| ECG          | No known CAD     | All                         | 29                | 3,392           | 62% (55%–68%)                | 68% (63%–73%)                |
|              |                  | Good                        | 10                | 1,410           | 70% (58%–79%)                | 62% (53%–69%)                |
|              | Mixed population | All                         | 41                | 4,879           | 61% (54%–67%)                | 65% (58%–72%)                |
|              |                  | Good                        | 13                | 1,679           | 65% (52%–76%)                | 60% (52%–68%)                |
| ECHO         | No known CAD     | All                         | 14                | 1,286           | 79% (74%–83%)                | 83% (74%–89%)                |
|              |                  | Good                        | 5                 | 561             | 79% (69%–87%)                | 85% (68%–94%)                |
|              | Mixed population | All                         | 22                | 1,873           | 78% (73%–83%)                | 86% (79%–91%)                |
|              |                  | Good                        | 8                 | 807             | 77% (65%–85%)                | 89% (76%–95%)                |
| SPECT        | No known CAD     | All                         | 14                | 1,000           | 81% (76%–86%)                | 78% (69%–84%)                |
|              |                  | Good                        | 4                 | 394             | 83% (52%–95%)                | 72% (37%–92%)                |
|              | Mixed population | All                         | 30                | 2,146           | 82% (77%–87%)                | 81% (74%–86%)                |
|              |                  | Good                        | 10                | 982             | 82% (72%–88%)                | 79% (66%–87%)                |
| CMR          | No known CAD     | All                         | 5                 | 501             | 72% (55%–85%)                | 84% (69%–93%)                |
|              |                  | Good                        | 5                 | 501             | 72% (55%–85%)                | 84% (69%–93%)                |
|              | Mixed population | All                         | 6                 | 778             | 78% (61%–89%)                | 84% (74%–90%)                |
|              |                  | Good                        | 5                 | 610             | 76% (55%–89%)                | 84% (72%–91%)                |
| Coronary CTA | No known CAD     | All                         | 5                 | 474             | 93% (69%–99%)                | 77% (54%–91%)                |
|              |                  | Good                        | 3                 | 124             | 85% (26%–99%)                | 73% (17%–97%)                |
|              | Mixed population | All                         | 8                 | 690             | 94% (81%–98%)                | 87% (68%–96%)                |
|              |                  | Good                        | 4                 | 201             | 83% (58%–94%)                | 77% (40%–94%)                |

Abbreviations: CAD = coronary artery disease; CI = confidence interval; CMR = cardiac magnetic resonance; CTA = computed tomography angiography; ECG = exercise/stress electrocardiogram; ECHO = echocardiogram; SPECT = single proton emission computed tomography



**Figure C. Summary of accuracy of NITs compared with coronary angiography for diagnosing CAD in women with no known CAD (all studies)**



**Table C. Summary of key findings**

| Key Question   | Strength of Evidence   | Conclusions   |
|--|--|---|
| <p><b>KQ 1: Diagnostic accuracy of NITs in women</b></p> | <p>ECG: High<br/>ECHO: High<br/>SPECT: High<br/>CMR: Low<br/>Coronary CTA: Low</p> | <p>94 studies described the diagnostic accuracy of NITs in comparison to another NIT or coronary angiography in women. Of these 94 studies, 78 studies included sufficient data to estimate the sensitivity and specificity of the NIT compared with coronary angiography.</p> <p>Summary from all studies with no known CAD:</p> <p>41 studies (13 good quality, 22 fair, 6 poor) of exercise ECG showed a summary sensitivity of 62% and specificity of 68%</p> <p>22 studies (8 good quality, 13 fair, 1 poor) of exercise/stress ECHO showed a summary sensitivity of 79% and specificity of 83%</p> <p>30 studies (10 good quality, 15 fair, 5 poor) of exercise/stress radionuclide perfusion imaging (SPECT, PET) showed a summary sensitivity of 81% and specificity of 78%</p> <p>6 studies (5 good quality, 1 fair) of CMR imaging showed a summary sensitivity of 72% and specificity of 84%</p> <p>8 studies (4 good quality, 4 fair) of coronary CTA showed a summary sensitivity of 93% and specificity 77%</p> <p>Overall, within a given modality, the summary sensitivities and specificities were similar for both types of populations (unknown CAD and mixed known and no known CAD) and for all studies when compared with good-quality studies. For the newer technologies (i.e., coronary CTA and CMR), more studies in women would be needed to support these findings since the 95 % CIs were quite wide.</p> <p>In testing for a statistically significant difference between the diagnostic accuracy of testing modalities in women, our analyses determined that for women with no previously known CAD, there were differences between the performance of the available modalities (<math>p &lt; 0.001</math>). The sensitivity of ECHO and SPECT was significantly higher than that of ECG. Specificity of ECG was less than that of CMR (borderline) and of ECHO. In the subset of studies that were good-quality and where there was no known CAD in the included population, our analyses again demonstrated differences between performance of tests (<math>p = 0.006</math>) with the specificity of ECG being less than that of CMR and ECHO.</p> |

**Table C. Summary of key findings (continued)**

| Key Question  | Strength of Evidence | Conclusions   |
|---|----------------------|---|
| <b>KQ 1 (continued)</b>   |                      | Sensitivity analyses exploring mixed populations of women with known and no known CAD showed no statistically significant difference in the sensitivities and specificities from our primary analysis. An analysis exploring the prevalence of CAD across the different NIT modality studies also showed no statistically significant difference. In addition, there were very few studies (1 SPECT, 1 ECHO, and 3 ECG) that did not complete a coronary angiography in all patients who underwent the NIT; therefore the results are minimized for verification bias. Finally we found no evidence of publication bias across the different modalities in our 4 populations of interest (studies of women with no known CAD, good quality studies of women with no known CAD, studies of women from mixed populations, and good quality studies of women from mixed populations).  |
| <b>KQ 2: Predictors of diagnostic accuracy in women</b>                           | Insufficient         | 11 studies (4 good quality, 5 fair, 2 poor) described diagnostic accuracy, and 9 of these examined predictors of diagnostic accuracy of different NITs in women.<br>Summary:<br>The predictors assessed included (1) postmenopausal women ages 55 to 64 (1 study), (2) race/ethnicity (2 studies), (3) heart size (4 studies), (4) pretest probability (3 studies), and (5) use of beta blocker medications (1 study).<br>We identified no studies examining the influence of age alone, functional status, or body size on diagnostic accuracy in women.<br>In terms of the NIT modality, we found four studies of stress ECHO, six studies of stress ECG, two studies of CMR, and four studies of SPECT that reported these predictors.<br>Insufficient evidence was available to draw definitive conclusions about predictors given the small number of studies for each predictor and for each modality, as well as the combination of predictor by modality. |
| <b>KQ 3: Improving risk stratification, decisionmaking, and outcomes in women</b> | Insufficient         | 13 studies (3 good quality, 9 fair, 1 poor) reported prognostic, outcome, or decisionmaking data comparing one NIT with another NIT or with coronary angiography in women with symptoms suspicious for CAD.<br>Summary:<br>We found 8 studies assessing risk stratification and prognostic information, 2 studies assessing decisionmaking for treatment options, and 4 studies that provided comparative clinical outcomes.<br>There were insufficient data to demonstrate that the use of specific NITs (compared with coronary angiography) routinely provided incremental risk stratification, prognostic information, or other meaningful information to improve decisionmaking and improve patient outcomes.<br>Most findings reported in the literature would require significant confirmation and replication in larger studies with women.   |

**Table C. Summary of key findings (continued)**

| <b>Key Question</b>          | <b>Strength of Evidence</b> | <b>Conclusions</b>   |
|------------------------------|-----------------------------|--|
| <b>KQ 4: Safety concerns</b> | Insufficient                | <p>13 studies (9 good quality, 4 fair) reported data pertinent to safety concerns or risks associated with the use of NITs to diagnose CAD in women with symptoms suspicious for CAD.</p> <p>Summary:</p> <p>Safety data were reported on the following modalities: (1) stress ECG (4 studies), (2) ECHO (6 studies), (3) SPECT (3 studies), (4) CMR (2 studies), and (5) coronary CTA (4 studies).</p> <p>Data specific to women on access site complications, contrast agent-induced nephropathy, nephrogenic systemic fibrosis, or anaphylaxis associated with NITs were not reported in any of the studies included in this report.</p> <p>Other than higher mean effective radiation doses for coronary CTA studies for women compared with men (from 3 out of 4 studies reporting radiation exposure levels), the extant literature does not provide sufficient evidence to conclude whether safety concerns, risks, or radiation exposure associated with different NITs to diagnose CAD in patients with suspected CAD differ significantly between women and men.</p> |

## Discussion

In summary, the findings of this comparative effectiveness review provide evidence for the accuracy of exercise/stress ECG, ECHO, SPECT, CMR, and coronary CTA compared with coronary angiography used for diagnosing CAD in women. The diagnostic accuracy appears to be consistent over time except for the sensitivity of CMR, which appears to be increasing over time (although the large confidence intervals reflect the underlying uncertainty in this measure). We are confident that the summary statistics for ECG, ECHO, and SPECT are robust and unlikely to change with the addition of new studies based on both the number of good-quality trials comparing these modalities with coronary angiography and the tight confidence intervals. More good-quality studies comparing CMR or coronary CTA with coronary angiography in the no-known CAD population and reporting sex-based results are needed to strengthen the summary statistics for those modalities.

Decisions around performing tests (either noninvasive or invasive) in patients with symptoms suspicious for CAD revolve around first understanding the pretest probability and testing/action thresholds for patients from the AHA/ACC stable angina guidelines and appropriate use criteria for the various NIT modalities.<sup>25-28</sup> Specifically, clinicians faced with patients who have a guideline-defined low-to-intermediate pretest probability of CAD may decide to obtain a noninvasive test, ideally with a high negative predictive value in this population and low risk of adverse events, in order to “rule out” disease. These may be patients with atypical chest pain (e.g., reflux or musculoskeletal disease) who are concerned about a heart problem and who require reassurance that their symptoms are not cardiac in origin. In contrast, in patients with high pretest probability of CAD (greater than 90 percent chance), a test with very high positive predictive value in this population and potentially more risk may be chosen since the disease of interest is thought to be present; in these cases, invasive angiography—the gold standard—is recommended by the current clinical practice guidelines. Finally, it is the spectrum of intermediate probability between 10 and 90 percent for which the clinicians must choose noninvasive tests that provide the right balance of sensitivity, specificity, and clinical risk to warrant testing. The choice of NIT may differ by clinician preference, availability, or setting (outpatient vs. chest pain unit of an emergency department).

It is in this context that the findings of this report on the effectiveness of NITs in women must be considered. First, women are thought to be at lower pretest probability of CAD when evaluated in comparison with men of the same age. When comorbidities or risk factors are taken into account, the pretest likelihood increases with a higher number of comorbidities. Second, women susceptible to some of the adverse effects of testing may have poor test performance or have higher rates of complication from invasive arterial access. Third, because of body shape and limited functional capacity, women may not obtain the same test performance that men do from noninvasive testing. Finally, because of the lack of full representation of women across the spectrum of disease, the available literature may not provide data on performance at the ends of the probability spectrum. Spectrum bias may be present since the studies we evaluated had potentially varied populations and varied disease definitions.

While readers may assume that requiring coronary angiography as the comparator would bias this report toward a higher risk CAD population, we found that the mean CAD prevalence ranged from 0.26 to 0.44; thus there was a broad spectrum of CAD prevalence in these studies. In fact, the range of CAD prevalence in this review is similar to a recent analysis of a large administrative database of patients referred for coronary angiography in which the prevalence of

significant obstructive disease was 38 to 40 percent.<sup>29</sup> The patient population that does not require coronary angiography can be characterized as having symptoms with low suspicion for CAD or pretest probability of less than 10 percent (note that all included studies enrolled patients with “suspected CAD”). Thus, results from this review would not apply to patients with low pretest probability of disease (e.g., gastroesophageal reflux, musculoskeletal pain, or panic attacks) where an NIT may be performed for clinical reassurance that their symptoms are noncardiac in origin.

In general, because there are few patients with high pretest probability, most clinicians would prefer to have patients undergo one NIT prior to determining a treatment choice or referral to coronary angiography. More than one NIT test is often required when the initial test results are equivocal. Our review did not identify studies that discussed the order in which different NITs were used for evaluating CAD. In fact, multiple testing or layered-testing strategies are areas where significant research is needed.

The current data suggest that NITs with higher sensitivity include coronary CTA and SPECT, and stress ECHO may represent an NIT with higher specificity. Stress CMR shows emerging data that may be in the upper range for both sensitivity and specificity. Additionally, the findings also demonstrate that NIT performance in women is not as good as in men, likely due to the reasons addressed above. The accuracy may also be location or operator dependent, and thus the results of published studies conducted at highly specialized centers may not uniformly apply to those seen in routine practice. Choice of NIT—and whether to use exercise or pharmacological stress imaging—may be influenced by functional capacity, which tends to be lower in women compared with men. Of note, the accuracy data for NIT modalities in men appeared a little higher than expected given previous meta-analyses of diagnostic accuracy data in the total population, which is likely because the published literature combined the accuracy data for men and women. Taken in context, these findings give support to the current ACC/AHA recommendations and studies on noninvasive testing in women.

Women are more likely than men to have false-positive stress tests; i.e., abnormal stress imaging with nonobstructive CAD on coronary angiography. In fact, up to 9 percent of women presenting with acute coronary syndrome will not have obstructive CAD when they undergo coronary angiography for potential PCI.<sup>30</sup> Some experts suggest that these phenomena are due to the presence of microvascular obstruction, the incidence of which is hard to determine since there is no clear diagnostic test used to establish the diagnosis.

Currently, there is debate on whether NITs that measure heart function abnormalities (ECG abnormalities, wall motion abnormality, ischemia), including exercise ECG, stress ECHO, and cardiac nuclear imaging, are equivalent or inferior to NITs that measure anatomic abnormalities (detection of CAD) by CMR or coronary CTA. Will knowing the coronary anatomy (nonobstructive or obstructive) in symptomatic patients lead to better implementation of secondary measures—control of blood pressure, diabetes, and hyperlipidemia—to reduce future cardiac events? Or is it more important to intervene with medications and/or revascularization when ischemia is present? Though this review does not answer these important questions, we describe this evidence gap in the Future Research section.

## **Limitations of This Review**

Despite identifying 104 studies (110 articles) that met the inclusion criteria, this systematic review has several limitations. First, our search focused on comparator studies of the various NITs with a gold standard of coronary angiography for establishing the diagnosis of CAD in

symptomatic patients. While this focus was adequate for identifying studies to assess the diagnostic accuracy of the NIT modalities in women, we found very few comparative studies that reported the influence of clinical characteristics or patient demographics on diagnostic accuracy. Few comparative studies (NIT vs. coronary angiography, or NIT vs. NIT) provided information on incremental risk stratification, prognostic information, or meaningful information regarding decisionmaking, and few reported the significant risks in women. Study results on these issues were reported for the total patient population and did not separate the effects by sex. Many of the included studies were single-sex (women) studies and limited our ability to fully evaluate sex differences. Also, by focusing on symptomatic patients, this report did not review the use of coronary artery calcium scoring for asymptomatic, high-risk populations.

We are aware that there are several noncomparator studies of each of the NIT modalities that address these issues in women since routine clinical care does not require two NIT modalities or an NIT modality plus coronary angiography for the diagnostic workup of suspected CAD. Given the focus on comparative effectiveness, we did not include these noncomparator studies in our review. By focusing the review on comparative studies, however, we are reducing the bias that is inherent in noncomparative studies. Noncomparative studies have selection, spectrum, and intervention biases for the following reasons: The choice of NIT is determined by the treating provider; only a subset of patients with indeterminate or positive results are referred for further NIT testing or coronary angiography; and the clinical outcomes may be influenced by the medical treatments or revascularization options that are offered. Second, the sample size and low representation of women in most of the comparator studies may affect the authors' ability to analyze the results by sex, therefore reducing the number of studies reporting these findings separately. Third, most studies lacked long-term followup of the patient population, which affected our ability to find studies that reported prognostic information on how the different NITs influenced clinical outcomes. Finally, our summary of the harms and risks of NITs is limited by the lack of disclosure of periprocedural and postprocedural complications in most of the studies.

## Conclusions

This systematic review has provided evidence for the summary sensitivities and specificities of exercise/stress ECG, ECHO, SPECT, CMR, and coronary CTA compared with coronary angiography in women. There was limited or insufficient evidence on the influence of clinical and demographic factors on comparative diagnostic accuracy, risk stratification, prognostic information, treatment decisions, clinical outcomes, and harms from different NITs specifically in women. Modifying the search criteria to include noncomparator studies of NIT modalities may increase the number of studies that address this limitation.

## Future Research

This comprehensive review of the comparative effectiveness of NIT modalities for diagnosing women with suspected CAD identified numerous gaps in evidence that would be suitable for future research and for improving the reporting of findings of NIT studies in the published literature.

**Randomized trials comparing functional versus anatomic modalities.** Almost all the studies reviewed were prospective observational studies where patients already scheduled for coronary angiography also underwent one or two NIT modalities to assess the diagnostic accuracy of the NITs. In routine clinical practice, clinicians order one type of NIT modality based on a patient's ability to exercise, test availability, and clinician preference. Exercise ECG,

stress ECHO, and nuclear imaging all measure functional parameters to assess for ischemia and obstructive CAD. Newer technologies such as coronary CTA and CMR offer clinicians the ability to evaluate anatomic parameters to assess both nonobstructive and obstructive CAD. A comparison of a functional testing strategy to an anatomic testing strategy for patients with symptomatic chest pain is currently being done in two large clinical trials (PROMISE [NCT001174550] and RESCUE [NCT01262625]). The information from these clinical trials could inform how the choice of an NIT modality affects prognosis, treatment decisions, and clinical outcomes.

**Studies assessing outcomes beyond diagnostic accuracy.** Our review found very few *comparative* NIT studies that looked at risk stratification, prognostic information, treatment decisions, and clinical outcomes. Future studies, whether observational or controlled clinical trials, should have long-term followup of patient cohorts to assess these factors. This is important because a positive NIT result could lead to further testing to establish the diagnosis of CAD as well as lead to more attention to secondary prevention for CAD. As stated previously, multiple testing or layered-testing strategies, plus the influence on risk-factor modification (e.g., medication prescriptions and adherence), are areas where significant research is needed.

**Studies of sufficient sample size and representation of women.** Many studies assessing the comparative diagnostic accuracy of an NIT modality with another NIT modality or with coronary angiography did not present a sample size calculation for the numbers needed per group. In addition, after excluding the women-only studies, the studies with both sexes had low representation of women. In order to assess sex differences in NIT diagnostic accuracy or the impact on clinical outcomes, a sufficient sample size is required to have adequate statistical power for subgroup analyses.

**Reporting sex and CAD population subgroups separately.** From 1,662 citations, we excluded 1,376 (83 percent) for not reporting data on women and 615 (37 percent) for looking only at a population with known CAD. Since publication of the AHRQ report on the use of NITs in women,<sup>6,7</sup> there has been an increase in the number of studies reporting sex-based differences. We encourage more reporting of women's results as well as separating the results from no known CAD and known CAD populations. One challenge we encountered in this review was that the primary data representing the numbers of TP, TN, FP, and FN were not presented in most studies and often needed to be back-calculated based on reported sensitivities and specificities and underlying disease prevalence for our quantitative synthesis. It would aid future comparisons of modalities if study authors were to report the primary data for women and men separately either within the article itself or within an online supplementary appendix.

**Assessing clinical and demographic factors that influence diagnostic accuracy.** Clinicians are taught that clinical factors such as weight, heart size, functional status, race/ethnicity, sex, age, and menopausal status can influence the diagnostic accuracy of various NIT modalities. However, we found very few comparative studies that looked at the impact of these clinical and demographic factors on the sensitivity and specificity of NIT results. More evidence about predictors affecting diagnostic cardiac testing is needed to support or dispel these long-held notions. Additional studies of the NIT modalities to assess differing symptomatology and timing at presentation, racial differences, various risk profiles, and different settings (outpatient, inpatient, emergency room) would be help to build the evidence base needed for clinical decisionmaking.

**Reporting of risk, harms, and/or safety outcomes.** Diagnostic procedures to screen for heart disease can result in harmful clinical events (nephropathy, radiation exposure, access site



complications). Systematic reporting of adverse events in publications—in total and by sex—should continue to clarify which NIT modalities are safe after they are approved for use in clinical practice.

## Glossary

|       |  |
|-------|--|
| AHA   | American Heart Association                 |
| ACC   | American College of Cardiology             |
| CAD   | coronary artery disease                    |
| CMR   | cardiac magnetic resonance imaging         |
| CTA   | coronary computed tomography angiography   |
| ECG   | electrocardiogram, electrocardiography     |
| ECHO  | echocardiogram, echocardiography           |
| KQ    | Key Question                               |
| MeSH  | Medical Subject Heading                    |
| NIT   | noninvasive technology                     |
| NSF   | nephrogenic systemic fibrosis              |
| PET   | positron emission tomography               |
| MI    | myocardial infarction                      |
| SPECT | single proton emission computed tomography |
| TEP   | Technical Expert Panel                     |

## References

1. Mieres JH, Shaw LJ, Arai A., et al. Role of noninvasive testing in the clinical evaluation of women with suspected coronary artery disease: Consensus statement from the Cardiac Imaging Committee, Council on Clinical Cardiology, and the Cardiovascular Imaging and Intervention Committee, Council on Cardiovascular Radiology and Intervention, American Heart Association. *Circulation* 2005;111(5):682-96. PMID: 15687114
2. Lloyd-Jones D, Adams RJ, Brown TM,, et al. Heart disease and stroke statistics—2010 update: a report from the American Heart Association. *Circulation* 2010;121(7):e46-e215. PMID: 20019324
3. Stangl V, Witzel V, Baumann G., et al. Current diagnostic concepts to detect coronary artery disease in women. *European Heart Journal* 2008;29(6):707-17. PMID: 18272503
4. Matchar DB, Mark DB, Patel MR., et al. Non-invasive imaging for coronary artery disease. Technology Assessment. Rockville, MD: Agency for Healthcare Research and Quality, October 2006. <http://www.cms.gov/determinationprocess/downloads/id34TA.pdf>. Accessed December 7, 2010.
5. Agency for Healthcare Research and Quality. Methods Guide for Effectiveness and Comparative Effectiveness Reviews. Rockville, MD: Agency for Healthcare Research and Quality. <http://www.effectivehealthcare.ahrq.gov/index.cfm/search-for-guides-reviews-and-reports/?pageaction=displayproduct&productid=318>. Accessed August 22, 2011.

6. Grady D, Chaput L, Kristof M. Results of Systematic Review of Research on Diagnosis and Treatment of Coronary Heart Disease in Women. Evidence Report/Technology Assessment No. 80. (Prepared by the University of California, San Francisco-Stanford Evidence-based Practice Center under Contract No 290-97-0013.) AHRQ Publication No. 03-0035. Rockville,MD: Agency for Healthcare Research and Quality. May 2003. <http://archive.ahrq.gov/downloads/pub/evidence/pdf/chdwom/chdwom.pdf>. Accessed July 14, 2011.
7. Grady D, Chaput L, Kristof M. Diagnosis and Treatment of Coronary Heart Disease in Women: Systematic Reviews of Evidence on Selected Topics. Evidence Report/Technology Assessment No. 81. (Prepared by the University of California, San Francisco-Stanford Evidence-based Practice Center under Contract No 290-97-0013.) AHRQ Publication No. 03-E037. Rockville, MD: Agency for Healthcare Research and Quality. May 2003. <http://www.ahrq.gov/downloads/pub/evidence/pdf/chdwomtop/chdwomtop.pdf>. Accessed July 14, 2011.
8. Whiting P, Rutjes AW, Reitsma JB,, et al. The development of QUADAS: a tool for the quality assessment of studies of diagnostic accuracy included in systematic reviews. *BMC Med Res Methodol* 2003;3:25. PMID: 14606960
9. Atkins D, Chang SM, Gartlehner G,, et al. Assessing applicability when comparing medical interventions: AHRQ and the Effective Health Care Program. *J Clin Epidemiol* 2011;64(11):1198-207. PMID: 21463926
10. Leeftang MM, Deeks JJ, Gatsonis C,, et al. Systematic reviews of diagnostic test accuracy. *Ann Intern Med* 2008;149(12):889-97. PMID: 19075208
11. Blyth CR, Still HA. Binomial Confidence-Intervals. *Journal of the American Statistical Association* 1983;78(381):108-116. PMID: ISI:A1983QG01500024
12. Rutter CM, Gatsonis CA. Regression methods for meta-analysis of diagnostic test data. *Acad Radiol* 1995;2 Suppl 1:S48-56; discussion S65-7, S70-1 pas. PMID: 9419705
13. Arends LR, Hamza TH, van Houwelingen JC,, et al. Bivariate random effects meta-analysis of ROC curves. *Med Decis Making* 2008;28(5):621-38. PMID: 18591542
14. Agency for Healthcare Research and Quality. Methods Guide for Medical Test Reviews [posted November 2010]. Rockville, MD. <http://archive.ahrq.gov/downloads/pub/evidence/pdf/chdwomtop/chdwomtop.pdf>. Accessed August 22, 2011.
15. Owens DK, Lohr KN, Atkins D,, et al. AHRQ series paper 5: Grading the strength of a body of evidence when comparing medical interventions--Agency for Healthcare Research and Quality and the Effective Health-Care Program. *J Clin Epidemiol* 2010;63(5):513-23. PMID: 19595577
16. Dodi C, Cortigiani L, Masini M,, et al. The incremental prognostic value of pharmacological stress echo over exercise electrocardiography in women with chest pain of unknown origin. *Eur Heart J* 2001;22(2):145-52. PMID: 11161916
17. Sanfilippo AJ, Abdollah H, Knott TC,, et al. Stress echocardiography in the evaluation of women presenting with chest pain syndrome: a randomized, prospective comparison with electrocardiographic stress testing. *Can J Cardiol* 2005;21(5):405-12. PMID: 15861257
18. Shaw LJ, Mieres JH, Hendel RH,, et al. Comparative Effectiveness of Exercise Electrocardiography With or Without Myocardial Perfusion Single Photon Emission Computed Tomography in Women With Suspected Coronary Artery Disease: Results From the What Is the Optimal Method for Ischemia Evaluation in Women (WOMEN) Trial. *Circulation* 2011. PMID: 21844080
19. Siegler JC, Rehman S, Bhumireddy GP,, et al. The Accuracy of the Electrocardiogram during Exercise Stress Test Based on Heart Size. *PLoS One* 2011;6(8):e23044. PMID: 21857990

20. Doyle M, Fuisz A, Kortright E,, et al. The impact of myocardial flow reserve on the detection of coronary artery disease by perfusion imaging methods: an NHLBI WISE study. *J Cardiovasc Magn Reson* 2003;5(3):475-85. PMID: 12882078
21. Lehmkuhl HB, Siniawski H, Lehmkuhl E,, et al. Value and limitations of dobutamine stress echocardiography in women with suspected coronary artery disease. *Zeitschrift fur Herz-, Thorax- und Gefasschirurgie* 2007;21(6):250-258. PMID: 2008012568
22. Lu C, Lu F, Fragasso G,, et al. Comparison of exercise electrocardiography, technetium-99m sestamibi single photon emission computed tomography, and dobutamine and dipyridamole echocardiography for detection of coronary artery disease in hypertensive women. *Am J Cardiol* 2010;105(9):1254-60. PMID: 20403475
23. Miller TD, Roger VL, Milavetz JJ,, et al. Assessment of the exercise electrocardiogram in women versus men using tomographic myocardial perfusion imaging as the reference standard. *Am J Cardiol* 2001;87(7):868-73. PMID: 11274942
24. Raman SV, Donnally MR, McCarthy B. Dobutamine stress cardiac magnetic resonance imaging to detect myocardial ischemia in women. *Prev Cardiol* 2008;11(3):135-40. PMID: 18607148
25. Hendel RC, Patel MR, Kramer CM,, et al. ACCF/ACR/SCCT/SCMR/ASNC/NASCI/CAI/SIR 2006 appropriateness criteria for cardiac computed tomography and cardiac magnetic resonance imaging: a report of the American College of Cardiology Foundation Quality Strategic Directions Committee Appropriateness Criteria Working Group, American College of Radiology, Society of Cardiovascular Computed Tomography, Society for Cardiovascular Magnetic Resonance, American Society of Nuclear Cardiology, North American Society for Cardiac Imaging, Society for Cardiovascular Angiography and Interventions, and Society of Interventional Radiology. *J Am Coll Cardiol* 2006;48(7):1475-97. PMID: 17010819
26. Taylor AJ, Cerqueira M, Hodgson JM,, et al. ACCF/SCCT/ACR/AHA/ASE/ASNC/NASCI/SCAI/SCMR 2010 Appropriate Use Criteria for Cardiac Computed Tomography. A Report of the American College of Cardiology Foundation Appropriate Use Criteria Task Force, the Society of Cardiovascular Computed Tomography, the American College of Radiology, the American Heart Association, the American Society of Echocardiography, the American Society of Nuclear Cardiology, the North American Society for Cardiovascular Imaging, the Society for Cardiovascular Angiography and Interventions, and the Society for Cardiovascular Magnetic Resonance. *J Cardiovasc Comput Tomogr* 2010;4(6):407 e1-33. PMID: 21232696
27. Douglas PS, Garcia MJ, Haines DE,, et al. ACCF/ASE/AHA/ASNC/HFSA/HRS/SCAI/SCCM/SCCT/SCMR 2011 Appropriate Use Criteria for Echocardiography. A Report of the American College of Cardiology Foundation Appropriate Use Criteria Task Force, American Society of Echocardiography, American Heart Association, American Society of Nuclear Cardiology, Heart Failure Society of America, Heart Rhythm Society, Society for Cardiovascular Angiography and Interventions, Society of Critical Care Medicine, Society of Cardiovascular Computed Tomography, and Society for Cardiovascular Magnetic Resonance Endorsed by the American College of Chest Physicians. *J Am Coll Cardiol* 2011;57(9):1126-66. PMID: 21349406
28. Bonow RO, Douglas PS, Buxton AE,, et al. ACCF/AHA methodology for the development of quality measures for cardiovascular technology: a report of the American College of Cardiology Foundation/American Heart Association Task Force on Performance Measures. *Circulation* 2011;124(13):1483-502. PMID: 21875906
29. Patel MR, Peterson ED, Dai D,, et al. Low diagnostic yield of elective coronary angiography. [Erratum in: *N Engl J Med*. 2010 Jul 29;363(5):498.] *N Engl J Med* 2010;362(10):886-95. PMID: 20220183

30. Patel MR, Chen AY, Peterson ED, et al. Prevalence, predictors, and outcomes of patients with non-ST-segment elevation myocardial infarction and insignificant coronary artery disease: results from the Can Rapid risk stratification of Unstable angina patients Suppress ADverse outcomes with Early implementation of the ACC/AHA Guidelines (CRUSADE) initiative. *Am Heart J* 2006;152(4):641-7. PMID: 16996828

# Introduction

## Background

Cardiovascular disease is the leading cause of mortality for women in the United States.<sup>1</sup> Coronary heart disease—which includes coronary artery (or atherosclerotic) disease (CAD), myocardial infarction (MI), acute coronary syndromes, and angina—is the largest subset of this mortality.<sup>1</sup> According to the American Heart Association (AHA), approximately one in three female adults has some form of cardiovascular disease. Since 1984, the number of deaths attributed to cardiovascular disease in women has exceeded that in men, reaching 454,613 in 2005—more than deaths from all forms of cancer combined.<sup>2</sup> It is estimated that 8.1 million women alive today have a history of heart attack, angina pectoris (chest pain or discomfort caused by reduced blood supply to the heart muscle), or both, and experts predict that in 2010 alone an estimated 370,000 women will have a new or recurrent MI. Overall, women who have had an acute MI—particularly those older than 55 years of age—have a worse prognosis than men, with a greater recurrence of MI and higher mortality.<sup>1</sup> More women (5.5 million) than men (4.3 million) have angina in total numbers. Among women older than 20 years of age, non-Hispanic black women have the highest incidence of angina (6.7 percent) when compared with non-Hispanic whites (4.3 percent) and Mexican Americans (4.5 percent).<sup>2</sup> However, the prevalence of CAD in women with chest pain is about 50 percent, as compared with 80 percent in men, which complicates diagnosis in women.<sup>3</sup>

The AHA suggests there is evidence showing that women at risk for CAD are less often referred for the appropriate diagnostic test than are men.<sup>1</sup> Coronary anatomy and pathology have traditionally been defined and identified by invasive, catheter-based x-ray angiography, also referred to as coronary angiography.<sup>4</sup> In this invasive procedure, a catheter is inserted into the femoral, brachial, or radial artery and passed up through the aorta to directly engage the right and left coronary arteries; an iodinated contrast agent is then injected into each artery while digital x-ray images are recorded.<sup>4</sup> The major benefits of invasive coronary angiography over noninvasive techniques are that the use of a catheter makes it possible to see the coronary arteries with greater anatomic precision and resolution and to combine diagnosis and treatment in a single procedure. The limitations of the procedure include the invasive nature of the test and the limited data on the functional impact of a luminal obstruction. These limitations are generally considered to be minor when compared with the benefits of the procedure, and coronary angiography is now the reference (gold) standard for clinical care of patients who have chest pain suggestive of CAD.

Coronary angiography, however, is not risk-free.<sup>4</sup> Arterial bleeding can occur at the access site, and manipulation of the catheter within the aorta and coronary arteries may cause an atherosclerotic embolus that, in turn, could result in stroke or heart attack. Separation of material from the inner lining of the artery may also cause a blockage downstream of the catheter tip. The contrast agent used during the procedure to visualize the coronary arteries may cause anaphylaxis or renal impairment or injury, and there is radiation exposure during the digital x-ray imaging. Although it is a rare occurrence, the catheter can puncture an artery and cause internal bleeding.

Coronary angiography is generally indicated in patients who have chest pain and are at high risk for CAD. For intermediate-risk patients, clinicians have a wide range of noninvasive diagnostic modalities to choose from, with wide variability in reported sensitivities and specificities. Noninvasive technologies (NITs) are especially important options for patients who

have contraindications to invasive catheterization, or for those who would be put at higher risk for complications with invasive screening. Included in this group would be patients who have a higher risk of an embolic stroke because of extensive vascular disease in the aorta; those with endocarditis involving the aortic valve; and those who are at high risk for developing a pseudoaneurysm at the site of catheter insertion because of underlying vascular disease.<sup>4</sup>

## **Types of Noninvasive Technologies**

NITs can assess functional status (i.e., ischemia or no ischemia) or visualize anatomic abnormalities (i.e., no CAD, nonobstructive CAD, or obstructive CAD). Types of NITs include:

- Functional modalities:
  - Exercise/stress electrocardiography (ECG) exercise/stress or resting
  - Exercise/stress echocardiography (ECHO) with or without a contrast agent
  - Exercise/stress radionuclide myocardial perfusion imaging, including single proton emission computed tomography (SPECT) and positron emission tomography (PET)
- Anatomic modalities:
  - Stress myocardial perfusion and wall motion magnetic resonance imaging (CMR)
  - Coronary computed tomography angiography (coronary CTA)

The AHA and the American College of Cardiology (ACC) recommend that women with suspected CAD should be classified as either symptomatic or asymptomatic and further classified as being at low, intermediate, or high risk for the disease to guide the decision about which diagnostic test to use first.<sup>1</sup> In 2005, the AHA developed a consensus statement on the role of noninvasive testing in the clinical evaluation of women with suspected CAD. In this statement, the AHA recommended that women who are symptomatic and at intermediate to high risk of having CAD should undergo noninvasive diagnostic studies (i.e., exercise electrocardiography and cardiac imaging studies) and that those who are asymptomatic and at low risk of CAD should not undergo cardiac imaging studies.<sup>1</sup> The AHA consensus statement was a thorough synopsis of the extant literature regarding the diagnosis of CAD in women with expert-guided recommendations for the workup of symptomatic women but did not include a comparative effectiveness review of the accuracy of the various NIT modalities in women.

## **Functional Modalities**

### **Electrocardiographic Modalities**

Treadmill testing with exercise ECG is the oldest and most commonly used form of stress testing. It is widely available and has low initial costs. According to the joint AHA/ACC guidelines, women should undergo exercise testing if they have an intermediate risk of CAD on the basis of symptoms and risk factors.<sup>5</sup> Factors that are unique to women (such as hormonal factors) have been reported to induce ECG changes during exercise that diminish the accuracy of the test. For exercise ECG, a positive test is defined, at peak exercise, by a significant ST segment horizontal or downsloping depression (significant level may vary by study as  $\geq 1$  mm, 1 - 2 mm, or  $\geq 2$  mm ST). ECG changes alone may not provide adequate prognostication. Exercise ECG has been recognized in the literature as being less sensitive and specific for diagnosing obstructive CAD in women than in men. Additional factors may improve the accuracy of the exercise test, such as chronotropic and hemodynamic responses to exercise. Despite sex-specific limitations, existing ACC/AHA guidelines propose that evidence of sex-specific limitations is

insufficient to remove the stress exercise ECG test as the initial test for symptomatic women at intermediate risk for CAD who have normal resting ECG results and are capable of exercise.<sup>1,5</sup> The AHA asserts that integrating other parameters into exercise scores (e.g., the Duke Treadmill Score, the ST/heart rate index) may improve the predictive value in women and that a positive ECG result in women indicates more diagnostic tests are necessary.<sup>1</sup>

Another ECG-based test is the newly developed Multifunction Cardiogram<sup>®</sup> (Cardiac Analytics, Powell, OH). With this resting ECG technology, patients are tested while lying in a supine position. From the multifunction cardiogram machine, five ECG wires with electrodes are attached to the patient at the four standard limb-lead and precordial-lead V5 positions. An automatic simultaneous 2-lead (leads V5 and II) ECG sampling is recorded for 82 seconds with amplification and digitization, and the ECG data are then transmitted to a data center via an encrypted Internet connection. Results are then compiled into a report that can be reviewed on the multifunction cardiogram unit itself or on any computer that has a Web browser. At present, this device is not widely available.<sup>6</sup>

### **Echocardiographic Modalities**

Exercise/stress ECHO is another noninvasive method for diagnosing CAD that provides information on the presence of left ventricular systolic or diastolic dysfunction, valvular heart disease, and the extent of infarction and stress-induced ischemia (defined as new or worsening wall motion abnormalities). Exercise ECHO can be performed using a treadmill or upright bicycle. In patients who cannot exercise, dobutamine is the most commonly used pharmacological stress agent. Vasodilator stress ECHO uses dipyridamole or adenosine. Contrast agents available for stress ECHO include SonoVue<sup>®</sup>, Optison<sup>™</sup>, and Luminity<sup>™</sup>. For stress ECHO, a positive test is defined by the evidence of new wall motion abnormalities in a different segment of the heart that can occur at stress only or at rest and stress. The AHA asserts that exercise/stress ECHO provides significantly higher specificity and accuracy for diagnosing obstructive CAD in women than standard exercise ECG testing does. Exercise/stress ECHO is recommended for women who are symptomatic and are at intermediate to high risk of CAD (women with suspected CAD must also have abnormal results from resting ECG), and dobutamine stress ECHO is recommended for women with normal or abnormal ECG results who are incapable of exercise.<sup>1</sup> The significant advantages of stress ECHO over ECG are superior diagnostic performance, ability to localize areas of ischemia, and the option of performing stress testing on patients who are unable to exercise.<sup>3</sup> According to a recent review, the overall sensitivities for exercise/stress ECHO are reported to be slightly lower in women than in men, although the specificities appear to be comparable for both.<sup>3</sup>

### **Myocardial Perfusion Imaging Modalities**

Exercise/stress myocardial perfusion imaging, which includes SPECT, PET, and scintigraphy, is a nuclear-based technique that uses a combination of test elements to diagnose CAD. Of the imaging modalities, exercise SPECT, PET, and scintigraphy can be performed by using a treadmill or an upright bicycle. In patients who cannot exercise, the pharmacologic stress agents are dobutamine, adenosine, and dipyridamole. The radionuclides commonly used are technetium Tc 99m sestamibi (MIBI), thallous chloride TL-201 (thallium) and fluorodeoxyglucose.

SPECT is the most commonly performed stress imaging test in the United States, especially for men and women who are unable to exercise.<sup>1</sup> Recently, the use of stress PET has increased. Parameters included in this modality are perfusion defects, global and regional left ventricular function, and left ventricular volumes. For myocardial perfusion imaging studies, a positive test is one that demonstrates reversible ischemia, and different scores can be used. The most frequently used is the summed stress score, which is a semiquantitative index obtained by adding the individual score derived from the 17 or 20 segments analyzed and scored during the stress study. Each segment is scored on a 5-point scale (0 = normal, 1 = slight reduction of tracer uptake, 2 = moderate reduction of tracer uptake, 3 = severe reduction of uptake, 4 = absence of uptake). A summed stress score <4 is considered normal; 4 to 8, mildly abnormal; 9 to 13, moderately abnormal; and >13, severely abnormal. Another score is based on the analysis of extent and severity of stress perfusion defect in the different segments of the left ventricle. This modality has been found to have technical limitations in women, including false-positive results, because of breast attenuation and a small left ventricular chamber size; however, recent advances in nuclear imaging have improved its accuracy (i.e., reduced the breast artifact).<sup>1</sup>

SPECT imaging is recommended for symptomatic women with an intermediate to high risk of CAD in the AHA 2005 consensus statement for the role of NIT in women.<sup>1</sup> A higher prevalence of single-vessel CAD among women adversely affects the diagnostic accuracy of this modality (as well as the ECHO modality).<sup>3</sup>

## **Anatomic Modalities**

ECG, ECHO, and myocardial perfusion imaging techniques do not provide direct visualization of coronary artery anatomy. They evaluate cardiac electrical activity, wall motion, or perfusion at rest and under stress, and any abnormal findings are used to make inferences about the presence and severity of obstructive CAD and the need for invasive coronary artery imaging. Other anatomic modalities provide direct visualization of coronary anatomy similar to that of coronary angiography but without invasive catheterization. These include cardiac magnetic resonance imaging (CMR) and coronary computed tomography angiography (coronary CTA). For CMR, a positive test is defined by the evidence of perfusion defects (extent and severity) and of wall motion abnormalities (at rest and/or at stress) in different left ventricular segments. For coronary CTA, a significant stenosis is defined quantitatively as at least a 50-percent narrowing (stenosis) of the coronary artery lumen.

Recently, the AHA published a scientific statement on CMR and coronary CTA in which recommendations were made for the general population and were not specific to women.<sup>7</sup> The AHA states that both tests are suboptimal for patients with atrial fibrillation and other arrhythmias, and image quality may be further reduced by a high body mass index. Overall, the AHA concludes that the potential benefit of noninvasive coronary angiography is likely to be greatest for symptomatic patients who are at intermediate risk for CAD after initial risk stratification, including patients with equivocal stress test results. The AHA does not recommend that CMR or coronary CTA be used to screen for CAD in patients without symptoms; in particular, concerns about the radiation dose limit the use of coronary CTA in patients who have a very low pretest likelihood of coronary stenoses. At the same time, patients with a high pretest likelihood of coronary stenoses are likely to require intervention and invasive catheter angiography for definitive evaluation. The AHA asserts that the main advantages of coronary CTA, compared with CMR, are wider availability, higher spatial resolution, and more consistent, shorter examinations with better patient adherence. Advantages associated with CMR include the



lack of need for ionizing radiation and an iodinated contrast agent. However, it is not clear whether the diagnostic accuracy or the relative balance of benefits and harms associated with either of these techniques differs between men and women.<sup>3,8</sup>

## **Uncertainties Surrounding Noninvasive Diagnosis of CAD in Women**

Noninvasive diagnosis of CAD in women is particularly challenging for many reasons. Women with chest pain demonstrate a lower prevalence of CAD, and their symptoms are less predictive and more often atypical when compared with those of men. Additionally, women are often older at the time of initial diagnosis; therefore, age-related comorbidities limit their tolerance for exercise testing.<sup>3</sup> Thus, many factors affect the accuracy of diagnostic testing for CAD in women, including:

- Lower prevalence of CAD
- Higher prevalence of nonobstructive CAD (microvascular abnormalities, mitral valve prolapse)
- Less predictive symptomatology
- Limited exercise tolerance because of older age, obesity, and diabetes at initial diagnosis
- Different response to exercise than men
- Lower peak exercise values
- Lesser increase in the left ventricular ejection fraction
- Increase in cardiac output by enhancing end-diastolic volume
- Inappropriate catecholamine release
- Hormonal influences of estrogens mimicking a digitalis-like false-positive ECG response
- Anatomic differences affecting stress test results
- Female breast attenuation artifacts
- Smaller coronary artery size
- Smaller left ventricular chamber size
- Higher prevalence of single-vessel disease
- Poor left ventricular opacification on echocardiography

In addition to all the factors that may affect the accuracy of noninvasive testing in women, there is currently considerable variation in which tests are used and in which order. In the acute-care setting, patients are often referred for early invasive coronary angiography as the initial risk stratification test although lower risk patients may be evaluated first with noninvasive testing.<sup>4</sup> After undergoing coronary angiography, some patients may be referred for noninvasive stress testing to define the functional significance of a coronary stenosis (constriction or narrowing) that is borderline in severity or is located such that the risk of treatment is increased.<sup>4</sup> Some cardiovascular experts advocate for a diagnostic strategy that includes both anatomic information (from direct coronary imaging, traditionally performed by using catheter angiography) and functional information collected during exercise or pharmacological stress testing.<sup>4</sup> Currently, there is no NIT modality that achieves both of these objectives.<sup>4</sup>

## Relevance

The goals of the diagnostic workup for women who have symptoms suspicious for CAD are to identify CAD with optimal accuracy and establish the basis for instituting preventive and therapeutic interventions.<sup>1</sup> More effective diagnostic strategies are critical for women at risk of CAD because up to 40 percent of initial cardiac events are fatal.<sup>2</sup> The literature suggests that, when compared with men, women are initially diagnosed with more advanced CAD because of the lack of early recognition and management.<sup>3</sup> Therefore, a better understanding of how the accuracy of the many different noninvasive tests for CAD varies by sex could dramatically improve outcomes for many women.

Noninvasive testing of women for CAD also raises uncertainty for decisionmakers because invasive coronary angiography has non-negligible patient risk. In addition, it is costly, requiring expensive equipment and the time and skill of highly trained physicians and support staff.<sup>7</sup> Although the use of noninvasive modalities to minimize the need for invasive procedures offers the possibility of better patient outcomes at lower cost, the wide range of diagnostic modalities—each with advantages and disadvantages for their use—makes it difficult for clinicians, patients, and payers to determine which test is best or should be used or covered in a given clinical situation.

## Scope and Key Questions

The following Key Questions (KQs) were considered in this comparative effectiveness review:

- **KQ 1.** What is the accuracy of one NIT in diagnosing obstructive and nonobstructive CAD when compared with another NIT or with coronary angiography in women with symptoms suspicious for CAD?
  - Exercise ECG stress test, including resting ECG technology (e.g., multifunctional cardiogram)
  - Exercise/stress ECHO with or without a contrast agent
  - Exercise/stress radionuclide myocardial perfusion imaging, including SPECT and PET
  - CMR imaging
  - Coronary CTA
- **KQ 2.** What are the predictors of diagnostic accuracy (e.g., age, race/ethnicity, body size, heart size, menopausal status, functional status, stress modality) of different NITs in women?
- **KQ 3.** Is there evidence that the use of NITs (when compared with other NITs or with coronary angiography) in women improves:
  - **KQ 3a.** Risk stratification/prognostic information?
  - **KQ 3b.** Decisionmaking regarding treatment options (e.g., revascularization, optimal medical therapy)?
  - **KQ 3c.** Clinical outcomes (e.g., death, myocardial infarction, unstable angina, hospitalization, revascularization, angina relief, quality of life)?

- **KQ 4.** Are there significant safety concerns/risks (i.e., radiation exposure, access site complications, contrast agent-induced nephropathy, nephrogenic systemic fibrosis, anaphylaxis, arrhythmias) associated with the use of different NITs to diagnose CAD in women with symptoms suspicious for CAD?

# Methods

## Topic Development and Refinement

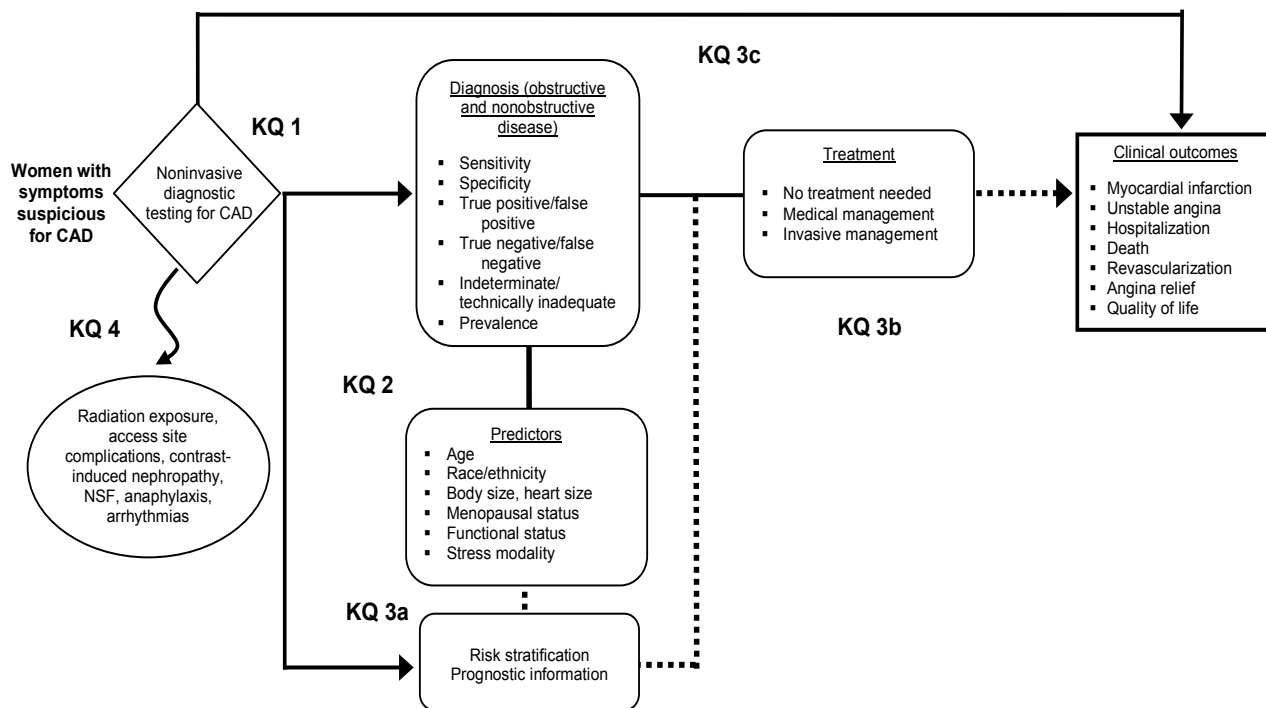
The Evidence-based Practice Center (EPC) followed AHRQ's recommended methodology, described in Methods Guide for Effectiveness and Comparative Effectiveness Reviews,<sup>9</sup> for literature search strategies, inclusion/exclusion of studies, abstract screening, data abstraction and management, assessment of methodological quality of individual studies, data synthesis, and grading of evidence for each Key Question (KQ).

During the topic refinement stage, we solicited input from Key Informants representing clinicians (cardiology, primary care, cardiac imaging), patients, scientific experts, and Federal agencies, to help define the KQs. The KQs were then posted for public comment for 30 days, and the comments received were considered in the development of the research protocol. We next convened a Technical Expert Panel (TEP), comprising clinical, content, and methodological experts, to provide input in defining populations, interventions, comparisons, or outcomes as well as identifying particular studies or databases to search. The Key Informants and members of the TEP were required to disclose any financial conflicts of interest greater than \$10,000 and any other relevant business or professional conflicts of interest. Any potential conflicts of interest were balanced or mitigated. Neither Key Informants nor members of the TEP did analysis of any kind and did not contribute to the writing of the report.

## Analytic Framework

Figure 1 shows the analytic framework for the systematic review of NITs for the diagnosis of CAD in women.

**Figure 1. Analytic framework**



Abbreviations: CAD = coronary artery disease, KQ = Key Question, NSF = nephrogenic systemic fibrosis

## Literature Search Strategy

### Sources Searched

We included studies published in English from January 1, 2000, through September 12, 2011. Search strategies were specific to each database in order to retrieve the articles most relevant to the KQs. Our search strategy used the National Library of Medicine’s medical subject headings (MeSH) keyword nomenclature developed for MEDLINE<sup>®</sup> and adapted for use in other databases. In consultation with our research librarians, we used PubMed<sup>®</sup>, Embase<sup>®</sup>, the Cochrane Database of Systematic Reviews, and the Cochrane Central Registry of Controlled Trials for our literature search. We also searched the grey literature of study registries and conference abstracts for relevant articles from completed trials. Grey literature databases included Clinicaltrials.gov; metaRegister of Controlled Trials; ClinicalStudyResults.org; WHO: International Clinical Trials Registry Platform Search Portal; CSA Conference Papers Index; and Scopus. The exact search strings used in our strategy are given in Appendix A. Reference lists of articles applicable to the relevant KQs of previous AHRQ reports on this topic<sup>10,11</sup> and from identified systematic reviews and meta-analyses were manually hand-searched and cross-referenced against our library, and additional manuscripts were retrieved. All citations were

imported into an electronic bibliographic database (EndNote<sup>®</sup> Version X4; Thomson Reuters, Philadelphia, PA).

## **Process for Study Selection**

### **Screening for Inclusion and Exclusion**

We developed a list of article inclusion and exclusion criteria for the KQs (Table 1). Using the prespecified inclusion and exclusion criteria, titles and abstracts were examined independently by two reviewers for potential relevance to the KQs. Articles included by any reviewer underwent full-text screening. At the full-text screening stage, two independent reviewers read each article to determine if it met eligibility criteria. At the full-text review stage, paired researchers independently reviewed the articles and indicated a decision to “include” or “exclude” the article for data abstraction. When the paired reviewers arrived at different decisions about whether to include or exclude an article, we reconciled the difference through a third-party arbitrator. Articles meeting our eligibility criteria were included for data abstraction. Relevant systematic review articles, meta-analyses, and methods articles were flagged for hand-searching and cross-referencing against the library of citations identified through electronic database searching.

**Table 1. Summary of inclusion and exclusion criteria**

| Study Characteristic | Inclusion Criteria  | Exclusion Criteria   |
|----------------------|---|--|
| Study design         | <p>An article was included if the following two criteria were met:</p> <ul style="list-style-type: none"> <li>• Original data or related methodology paper of an included article</li> <li>• Randomized controlled trial, prospective or retrospective observational study, or registry</li> </ul>  | <ul style="list-style-type: none"> <li>• Editorial</li> <li>• Letter to editor</li> <li>• Case series</li> <li>• Review article, meta-analysis, or methods paper of an excluded article</li> <li>• Not peer reviewed</li> </ul>  |
| Population           | <p>Study included adult women (age <math>\geq</math> 18 years of age) who present symptoms of symptoms suspicious for CAD (e.g., exertional dyspnea, shortness of breath, and/or angina) with or without a known diagnosis of CAD; data for women must be presented separately from data for men</p>  | <ul style="list-style-type: none"> <li>• All subjects were &lt; 18 years of age, or some subjects were under &lt; 18 but results were not broken down by age</li> <li>• No patients had symptomatic chest pain (i.e., an asymptomatic population), or some of the patients had symptomatic chest pain but results were not reported separately for this subgroup</li> <li>• All patients were known to have CAD and were not being tested for chest pain symptoms (e.g., postrevascularization testing to assess for persistent ischemia)</li> </ul> |
| Interventions        | <p>NITs for the diagnosis of obstructive and nonobstructive CAD included:</p> <ul style="list-style-type: none"> <li>• Exercise electrocardiogram stress test</li> <li>• Resting electrocardiogram technology</li> <li>• Exercise/stress echocardiography with or without a contrast agent</li> <li>• Exercise/stress radionuclide myocardial perfusion imaging, including single proton emission computed tomography and positron emission tomography</li> <li>• Stress myocardial perfusion and wall motion magnetic resonance imaging</li> <li>• Coronary computed tomography angiography</li> </ul> | <p>Coronary artery calcium scoring by electron beam computed tomography since this modality is often used to screen asymptomatic patients for CAD</p>  |
| Comparators          | <p>Another NIT or coronary angiography</p>  | <p>Study did not compare one NIT with another, or with coronary angiography</p>  |

**Table 1. Summary of inclusion and exclusion criteria (continued)**

| Study Characteristic  | Inclusion Criteria  | Exclusion Criteria   |
|-----------------------|---|--|
| Outcomes              | Primary outcome—accurate diagnosis of obstructive and nonobstructive CAD  | Outcomes not related to diagnostic accuracy for detecting CAD  |
|                       | KQ 1 patient-level outcomes: <ul style="list-style-type: none"> <li>• Sensitivity</li> <li>• Specificity</li> <li>• True positive, false negative, true negative, false positive</li> <li>• Indeterminate or technically inadequate results</li> <li>• Prevalence</li> </ul>  | <ul style="list-style-type: none"> <li>• Vessel-based outcomes</li> <li>• Outcomes of women not reported separately from total population</li> </ul> |
|                       | KQ 2 outcomes: Predictors of diagnostic accuracy—age, race/ethnicity, body size, heart size, menopausal status, functional status, stress modality  | Outcomes of women not reported separately from total population  |
|                       | KQ 3 outcomes: <ul style="list-style-type: none"> <li>• Risk stratification/prognostic information</li> <li>• Treatment —none, medical therapy, percutaneous coronary intervention, or coronary artery bypass surgery</li> <li>• Clinical outcomes—death, myocardial infarction, unstable angina, hospitalization, revascularization, angina relief, quality of life</li> </ul> | Outcomes of women not reported separately from total population  |
|                       | KQ 4 outcomes: Safety and adverse events—radiation exposure, access site complications, contrast agent-induced nephropathy, nephrogenic systemic fibrosis, anaphylaxis, arrhythmias—and how these events varied by demographic factors  | Outcomes of women not reported separately from total population  |
| Setting               | Inpatient or outpatient settings, primarily primary care and cardiology clinics   | None   |
| Publication languages | English only  | Given the high volume of English-language publications (including the majority of known important studies), non-English articles were excluded       |

Abbreviations: CAD = coronary artery disease; KQ = Key Question; NIT = noninvasive technology

## Data Extraction and Data Management

The investigative team created forms for abstracting the data elements for the KQs. The data abstraction forms were piloted by two members of the study team, and refinements to clarify the questions and collection of data were added after the first week of data abstraction. The investigators who piloted the forms were also the main data abstractors. Based on their clinical and methodological expertise, two researchers were assigned to abstract data from the eligible articles pertaining to the research questions. One researcher abstracted the data, and the second overread the article and the accompanying abstraction form to check for accuracy and completeness. Disagreements were resolved by consensus or by obtaining a third reviewer’s opinion if consensus was not reached by the first two researchers. Any data changes during overreading were noted in a comment field. The overall interrater reliability was 0.76.



To aid in both reproducibility and standardization of data collection, researchers received data abstraction instructions directly on each form created specifically for this project with the DistillerSR data synthesis software program (Evidence Partners Inc., Manotick, ON, Canada). We designed these forms to collect the data required to evaluate the specified eligibility criteria for inclusion in this review as well as to collect demographics and data needed to determine outcomes (intermediate outcomes, health outcomes, and safety outcomes). The safety outcomes were framed to help identify radiation exposure, contrast agent-induced nephropathy, nephrogenic systemic fibrosis, anaphylaxis, and arrhythmias, which are the more common adverse events resulting from use of the different NITs. The abstraction form templates were pilot tested with a sample of included articles to ensure that all relevant data elements were captured and that there was consistency and reproducibility between abstractors. Appendix B lists the elements used in the data abstraction forms.

Appendix C contains a bibliography of all studies included in this review, organized alphabetically by author. When appropriate, methods articles providing additional detail were considered when abstracting data for an included study. If a methods article was used as a source for information in the abstraction of a study, it was included in the review and is listed in the bibliography in Appendix C.

## **Individual Study Quality Assessment**

The studies included in this comparative effectiveness review were assessed on the basis of the quality of their reporting of relevant data. We evaluated the quality of individual studies using the approach described in AHRQ's Methods Guide for Effectiveness and Comparative Effectiveness Reviews (hereafter referred to as the Methods Guide).<sup>9</sup> To assess study quality, we (1) classified the study design, (2) applied predefined criteria for quality and critical appraisal, and (3) made a summary judgment of the study's quality. To evaluate methodological quality, we applied criteria for each study type that were derived from the core elements described in the Methods Guide<sup>9</sup> and from QUADAS,<sup>12</sup> a tool for the quality assessment of studies of diagnostic accuracy included in systematic reviews. To indicate the summary judgment of the quality of the individual studies, we used the summary ratings of Good, Fair, and Poor based on the study's adherence to well-accepted standard methodologies (such as QUADAS) and adequate reporting standards.

Grading was outcome specific; thus, a given study may have been graded of different quality for two individual outcomes reported within that study. Study design was considered when grading quality. Randomized controlled trials were graded as Good, Fair, or Poor. Observational studies were graded separately also as Good, Fair, or Poor.

We used data abstracted on the population studied, the intervention and comparator, the outcomes measured, settings, and timing of assessments to identify specific issues that may have limited the applicability of individual studies or a body of evidence as recommended in the Methods Guide.<sup>9,13</sup> We used these data to evaluate the applicability to clinical practice, paying special attention to study eligibility criteria, demographic features of the enrolled population in comparison with the target population, the intervention used in comparison with technologies currently in use, and clinical relevance and timing of the outcome measures. We summarized issues of applicability qualitatively. The quality grading and applicability were conducted in similar fashion to the rest of the data abstraction phase, with one investigator entering ratings, a second investigator overreading the entries, and disagreements resolved through consensus or a

third party. Appendix D summarizes our assessment of the quality and applicability for each included study as well as the assessed QUADAS quality scores for diagnostic accuracy.

## Data Synthesis

We summarized the primary literature by abstracting relevant continuous data (e.g., age, sensitivity, specificity, event rates) and categorical data (e.g., race/ethnicity, presence of CAD). Data for patients with no known diagnosis of CAD were collected and analyzed separately from data for mixed CAD populations that included patients with and without known CAD. We then determined the feasibility of completing a quantitative synthesis (i.e., summary receiver operating characteristic [SROC] curves for diagnostic accuracy or meta-analysis for other outcomes). The feasibility of a meta-analysis or SROC curve depended on the volume of relevant literature, the homogeneity of the studies in terms of the populations studied, the interventions included, or the outcomes assessed, and the completeness of the results reporting. For each SROC calculation, we ran separate analyses of the accuracy of each NIT modality compared with coronary angiography on the no-known CAD and mixed CAD populations using random-effects models to quantitatively synthesize the available evidence. In our primary analyses, we evaluated these performance characteristics in the population of women who had no previously known CAD. In secondary analyses, we explored a broader patient population by including those studies that had women from a mixed population of known and no known CAD. We also assessed the impact on our findings if, in each population, we restricted our analyses to those studies that were assessed to be good quality. We then compared the performance characteristics of the NIT modalities with each other using a generalized linear mixed model to assess for differences in summary sensitivity and specificity between the NIT modalities (ECG, ECHO, SPECT, CTA, CMR), as well as differences in disease state (no known and mixed CAD). In a final exploratory analysis, we evaluated the test performance of the modalities in women compared with men in a separate generalized linear mixed model with sex (women, men) as a covariate. We presented summary estimates and confidence intervals (CIs).

For synthesizing the accuracy data for studies included in our assessment of KQ 1, we used the following approach as advocated by Leeflang, et al.<sup>14</sup> This approach allows the paired nature of sensitivity and specificity and randomness between studies to be taken into account. The analyses are based on true positive (TP), false negative (FN), false positive (FP), and true negative (TN) frequencies abstracted from relevant publications. Estimated study specific sensitivity ( $TP/[TP+FN]$ ) and specificity ( $TN/[TN+FP]$ ) values are displayed in paired forest plots together with exact 95 % CIs.<sup>15</sup> The summary estimates of sensitivity and specificity resulted from random-effects modeling with two random effects, with each study being considered a random realization of the underlying true distribution. In the absence of covariates, both the bivariate random-effects model and the hierarchical model with random effects are mathematically equivalent.<sup>16</sup> We used the GLIMMIX procedure in the SAS statistical package (SAS Institute; Cary, NC) with maximum likelihood estimation. A binomial error model was used with the logit link. The unstructured covariance matrix of the two random effects was considered during the fitting process, and its Cholesky parameterization was used.

The fixed-effects estimates and their variance–covariance matrix provided (after reverse logit transformation) summary sensitivity and specificity values and a joint confidence region (dotted oval shape on figures) as well as separate CIs for summary sensitivity and specificity as presented on figures and forest plots in the report. There were several possible choices for the SROC curve resulting from the random-effects modeling.<sup>17</sup> We used the Rutter and Gatsonis<sup>18</sup>

SROC curve as described by Arends,, et al.,<sup>17</sup> and it is presented in figures as a solid line over the range of the available data.

In addition to our SROC curve estimation, we evaluated other available outcomes. Most outcomes that we analyzed in this comparative effectiveness report were binary or categorical, and so we summarized these outcomes by proportions. We summarized inherently continuous variables, such as age, by mean, median, and standard deviation.

We also evaluated the potential of verification bias and other limitations of our synthesized analyses based on the underlying clinical domain and diagnostic testing practices. For example, angiography is often administered only to a subset of patients who are undergoing diagnostic tests within a studied population. This subset of patients is not a completely random sample because angiography-based verification of disease is often driven by previous test results and/or other considerations. We explored the potential for publication bias across the different modalities in our four populations of interest (studies of women with no known CAD, good-quality studies of women with no known CAD, studies of women from mixed populations, and good-quality studies of women from mixed populations). Using methods advocated by Deeks,, et al.,<sup>19</sup> we computed for each study the diagnostic odds ratio (DOR) and the effective sample size (ESS). Subsequently we performed regression of natural logarithm of DOR against  $1/ESS^{1/2}$  weighing by ESS, and reported the p value for testing whether slope is equal to zero. A nonsignificant p value indicated no evidence for publication bias.

To explore additional sources of potential bias, we also recorded whether the diagnostic tests were interpreted in a blinded fashion; that is, without knowledge of results of other diagnostic tests or clinical history and risk factors, if such information was available in the reviewed studies.

## Grading the Body of Evidence

The strength of evidence for each KQ was assessed using the approach described in AHRQ's Methods Guide on Medical Test Reviews for grading the evidence related to the diagnostic accuracy of the NITs (KQ 1),<sup>20</sup> and the Methods Guide for grading the evidence related to the other KQs (KQs 2–4).<sup>9,21</sup> The outcomes used for grading the body of evidence for each KQ are outlined in Table 1. The evidence was evaluated using the four required domains: risk of bias (low, medium, or high), consistency (consistent, inconsistent, or unknown/not applicable), directness (direct or indirect), and precision (precise or imprecise). Additionally, when appropriate, the studies were evaluated for the presence of confounders that would diminish an observed effect, the strength of association (magnitude of effect), and publication bias. The strength of evidence was assigned an overall grade of High, Moderate, Low, or Insufficient according to the following four-level scale:

- High—High confidence that the evidence reflects the true effect. Further research is very unlikely to change our confidence in the estimate of effect.
- Moderate—Moderate confidence that the evidence reflects the true effect. Further research may change our confidence in the estimate of effect and may change the estimate.
- Low—Low confidence that the evidence reflects the true effect. Further research is likely to change the confidence in the estimate of effect and is likely to change the estimate.
- Insufficient—Evidence either is unavailable or does not permit estimation of effect.

## Peer Review and Public Commentary

The peer review process was our principal external quality-monitoring device. Nominations for peer reviewers were solicited from several sources, including the TEP and interested Federal agencies. The list of nominees was forwarded to AHRQ for vetting and approval. A list of reviewers submitting comments on this draft is included in the Preface of this report.

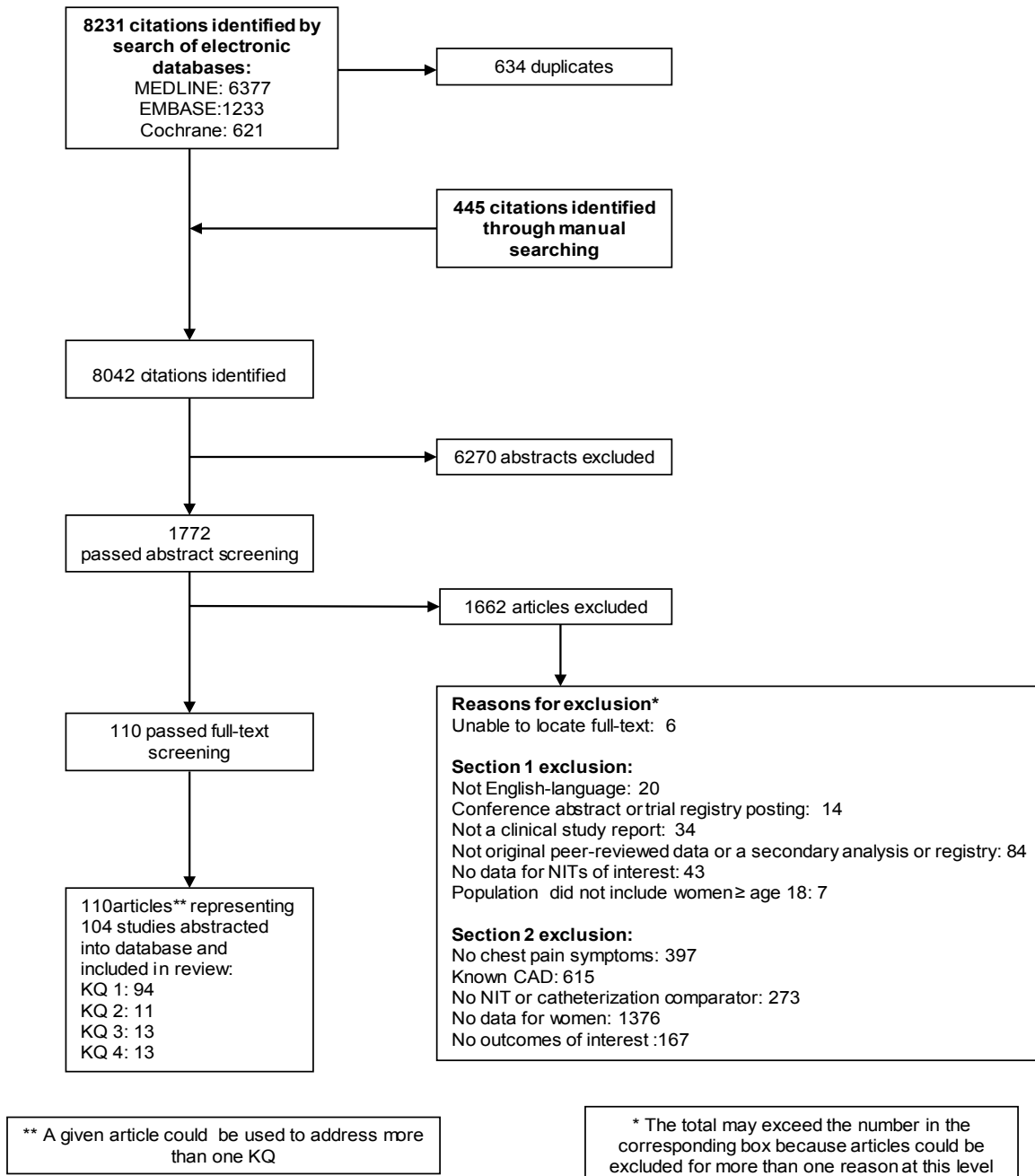
## Results

The flow of articles through the literature search and screening process is depicted in Figure 2. Of the 8,231 citations identified by our searches, 634 were duplicates. A manual search identified an additional 445 citations for a total of 8,042 citations. After applying inclusion/exclusion criteria at the title-and-abstract level, 1,772 full-text articles were retrieved and screened. Of these, 1,662 articles were excluded at the full-text screening stage. We excluded 1376 (83 percent) for not reporting data on women and 615 (37 percent) for looking only at a population with known CAD. (Note that an article may have been excluded for more than one reason.) The final set comprised 110 articles representing 104 studies.

Of the 104 studies, 1 was an RCT, 79 were prospective observational, and 24 were retrospective observational with study cohorts comprising individuals who presented for NIT testing and received diagnostic coronary angiography (100 studies) or another NIT modality only (4 studies). The four studies *without* coronary angiography compared ECHO with ECG<sup>22,23</sup> or ECG with SPECT.<sup>24,25</sup> Three of these studies were applicable to Key Question (KQ) 3,<sup>22-24</sup> and one was applicable to KQ 2.<sup>25</sup> Of the 94 studies included in the KQ 1 results, 5 reported NIT versus NIT comparisons in addition to coronary angiography.<sup>26-30</sup>

Appendix E provides a complete list of articles excluded at the full-text screening stage, with reasons for exclusion.

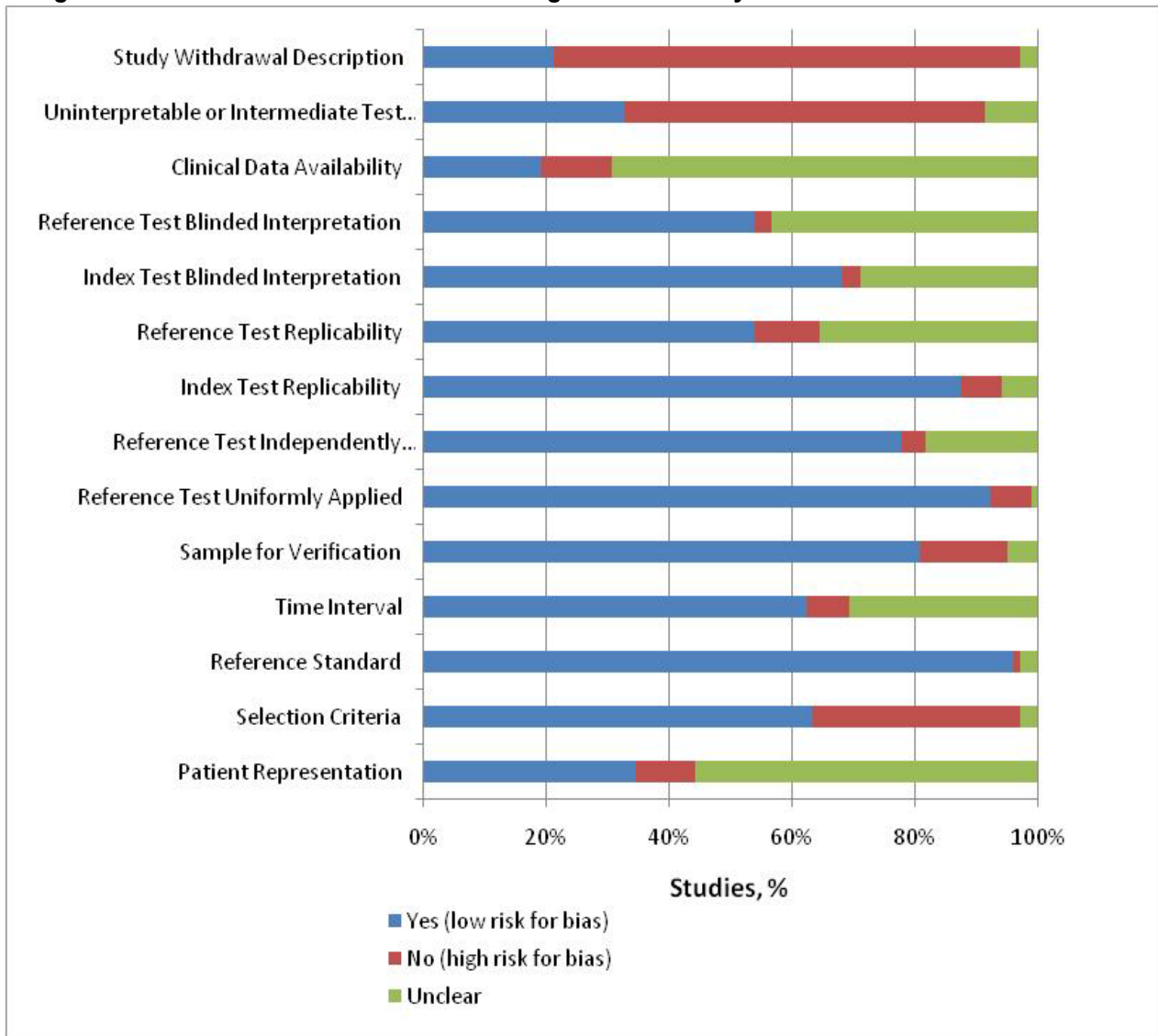
**Figure 2. Literature flow diagram**



Abbreviations: CAD = coronary artery disease; KQ = Key Question; NIT = noninvasive technology; SR = systematic review

A summary graph of the QUADAS ratings for all 104 studies is shown in Figure 3. (Refer to Table D-2 in Appendix D for a summary table of the QUADAS quality scores for diagnostic accuracy of the 104 studies included in this review.) A majority of studies uniformly applied a reference test (i.e., coronary angiography) that was independently performed, used blinded interpretation of the reference and index test, and had sufficient detail about the index test to allow for replicability. Many studies had a high risk of spectrum bias (i.e., patient representation of those who would receive the test in practice), a poor description of study withdrawals, unclear descriptions of clinical data available during test interpretation, and lacked descriptions of uninterpretable or intermediate test results. There is a possibility that sensitivity and specificity values may be biased because of subjects included in the studies that did not represent the spectrum of the population of interest; we explored the impact of the underlying prevalence of CAD in the population on our findings.

**Figure 3. QUADAS elements used to rate diagnostic accuracy**



## Key Question 1: Diagnostic Accuracy of NITs

KQ 1. What is the accuracy of one NIT in diagnosing obstructive and nonobstructive CAD when compared with another NIT or with coronary angiography in women with symptoms suspicious for CAD?

- Exercise ECG stress test, including resting ECG technology (e.g., multifunctional cardiogram)
- Exercise/stress ECHO with or without a contrast agent
- Exercise/stress radionuclide myocardial perfusion imaging, including SPECT and PET
- CMR imaging
- Coronary CTA

### Key Points

Individual study performance characteristics were evaluated for each testing modality, and summary receiver operating characteristic (SROC) curves were calculated. These analyses demonstrated:

- Overall, within a given testing modality, the summary sensitivities and specificities were similar for both types of populations (known and no known CAD) and for all studies when compared with good-quality studies.
- When accounting for only the good-quality studies, it appeared that the diagnostic accuracy of detecting CAD in women was better (in descending order) for coronary CTA, SPECT, ECHO, CMR, and ECG, although the strength of evidence varied markedly for different modalities.
- For the newer technologies (i.e., CMR and coronary CTA), more studies in women would be needed to support the point estimates given the wide confidence intervals (CIs) on the test performance.
- For women without previously known CAD, there were statistically significant differences between the performance of the available modalities ( $p < 0.001$ ). The sensitivity of ECHO and SPECT was significantly greater than that of ECG. Specificity of ECG was less than that of CMR (borderline) and of ECHO.
- In the subset of studies that were good-quality and where there was no known CAD in the included population, there were statistically significant differences between performance of tests ( $p = 0.006$ ), with the specificity of ECG being less than that of CMR and ECHO. Our ability to quantify the difference between test performance of the modalities between men and women was inhibited by the limited number of studies that reported both sexes separately in their analysis.



- In exploratory analysis of the difference between test performance in men and women, the ECG and coronary CTA modalities were both less sensitive and less specific in women than in men. The ECHO and SPECT modalities, although less sensitive, appeared to be more specific in women. The lower specificity of the ECG modality in women, however, is the only estimate that was determined to be a statistically significant difference.

## Detailed Synthesis

In KQ 1 we sought to determine the accuracy of each NIT modality in diagnosing obstructive and nonobstructive CAD when compared with coronary angiography in women with symptoms suspicious for CAD. For this analysis, we included 94 studies describing comparative diagnostic accuracy of NITs. Of these 94 studies, 78 studies included sufficient data to estimate the sensitivity and specificity of the NIT compared with coronary angiography. This included 41 studies examining exercise/stress ECG (13 good quality, 22 fair, 6 poor); 22 examining exercise/stress ECHO (8 good quality, 13 fair, 1 poor); 30 examining exercise/stress radionuclide myocardial perfusion imaging (e.g., SPECT or PET) (10 good quality, 15 fair, 5 poor); 6 examining CMR (5 good, 1 fair); and 8 examining coronary CTA (4 good quality, 4 fair).

For each testing modality, we used the individual performance characteristics to calculate an SROC curve and to estimate the summary sensitivity and specificity and CIs of the modality compared with coronary angiography. We present forest plots of the individual study estimates of sensitivity and specificity of each NIT for diagnosing CAD in women. Error bars in these plots represent 95 % CIs; the dashed vertical line represents the summary sensitivity and specificity for the included studies. The ROC curve illustrates the tradeoff between sensitivity and specificity since the threshold that defines a positive test result varies from the most stringent to the least stringent. Open circles represent individual study estimates of sensitivity and specificity. The black circle indicates the average sensitivity and specificity estimate of the study results, and the dashed circle represents the 95-percent confidence region around it. In our primary analyses, we evaluated these performance characteristics in the population of women who had no previously known CAD. In secondary analyses we explored a broader patient population including those studies that had women from a mixed population of known and no known CAD. We also assessed the impact on our findings if, in each population, we restricted our analyses to those studies that were assessed to be good quality. We then compared the performance characteristics of the NIT modalities with each other. In a final exploratory analysis, we evaluated the test performance of the modalities in women compared with men. All secondary analyses involved the use of separate generalized linear mixed models with covariates for disease state (no known versus mixed), NIT modality (ECG, ECHO, SPECT, CMR, CTA), or sex (women versus men).

## **ECG**

We identified 41 studies evaluating the accuracy of exercise/stress ECG compared with coronary angiography (Table 2).<sup>27-29,31-68</sup> This table lists those studies that focused purely on women with no known CAD and then follows with the additional studies that included a mixed population of known and no known CAD. Within these populations, good-quality studies are listed first, followed by those of fair and poor quality. Twenty-nine of the ECG studies reported accuracy data in women with no known CAD, and these are the studies used in our primary analysis.

In our secondary analysis, we evaluated the accuracy of exercise/stress ECG in diagnosing CAD in mixed populations of known and no known CAD. This analysis included 1 study that reported additional data for a mixed population<sup>61</sup> and an additional 12 studies that reported findings with mixed populations of known and no known CAD. Two of these studies evaluated the use of resting ECG,<sup>33,49</sup> and two studies<sup>35,50</sup> evaluated pharmacological stressed ECG. All other studies evaluated exercise/stress ECG.

### **Primary Analysis: Population of Women With No Known CAD**

The 29 studies represent findings on ECG use in 3,391 women (sample size ranging from 10 to 580 women). Of these studies, 10 were good quality, 15 were fair quality, and 4 were poor quality. Sensitivity varied from 32 to 91 percent, and specificity varied from 40 to 100 percent; the median sensitivity was 61 percent, and the median specificity was 68 percent. Figure 4 presents forest plots of the individual study estimates of sensitivity and specificity of ECG for diagnosing CAD in women with no known CAD. Error bars represent 95 percent CIs; the dashed vertical line represents the summary sensitivity and specificity for the included studies.

**Figure 4. Accuracy of ECG in women with no known CAD**

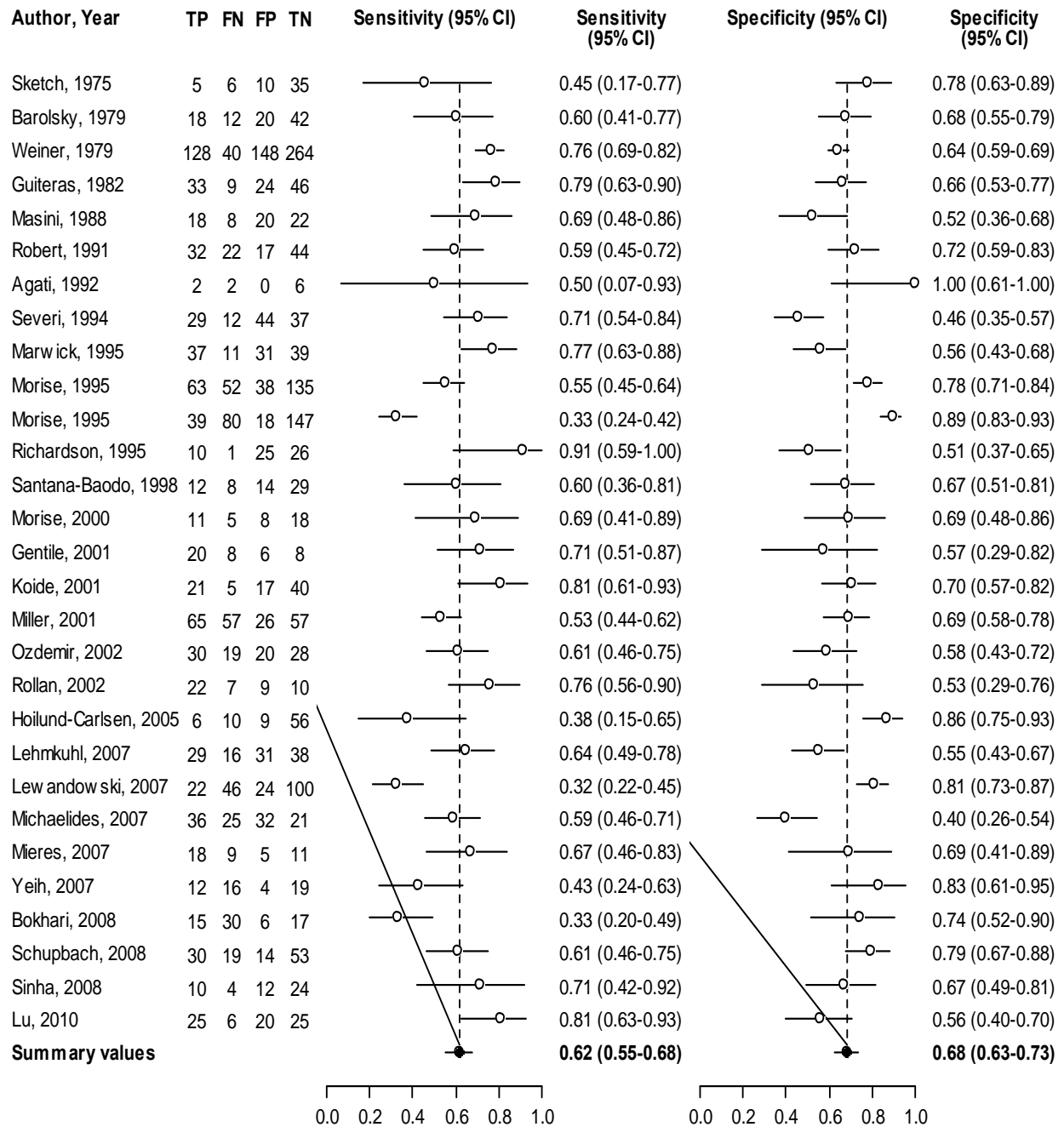
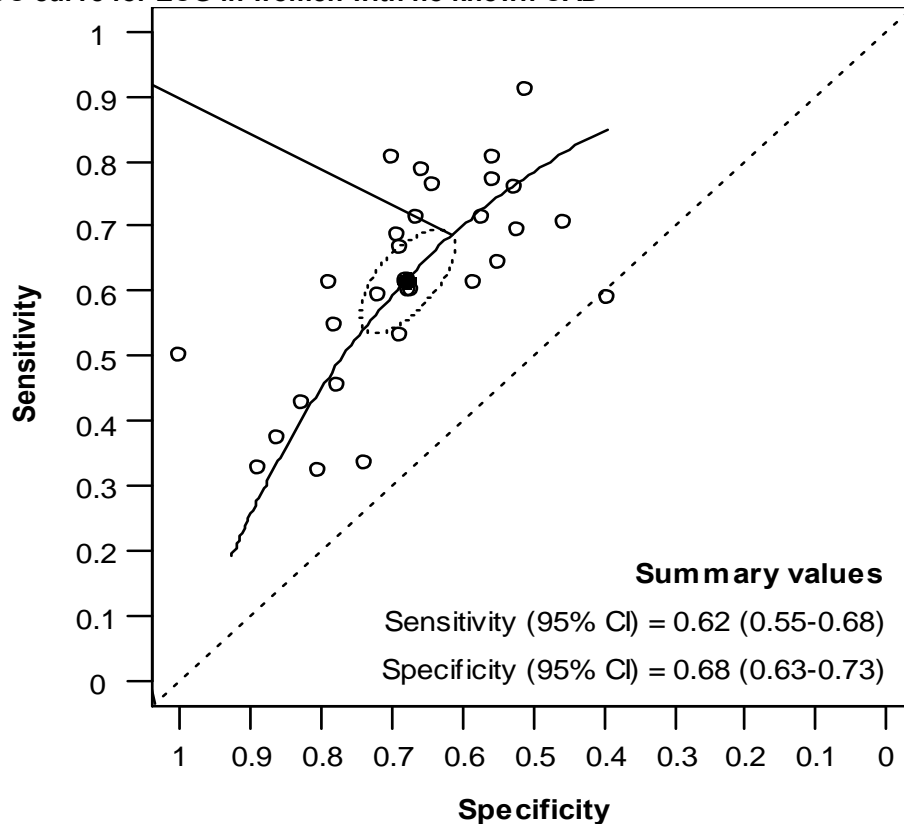


Figure 5 presents a summary receiver operating characteristic (SROC) curve with an average sensitivity of 62 percent (95% CI, 55 to 68 percent) and specificity of 68 percent (95% CI, 63 to 73 percent). The ROC curve illustrates the tradeoff between sensitivity and specificity since the threshold that defines a positive test result varies from the most stringent to the least stringent. Open circles represent individual study estimates of sensitivity and specificity. The black circle indicates the average sensitivity and specificity estimate of the study results, and the dashed circle represents the 95-percent confidence region around it.

**Figure 5. SROC curve for ECG in women with no known CAD**



The prevalence of CAD on coronary angiogram in these 29 studies ranged from 18 to 67 percent with a mean prevalence of 41 percent. In the individual studies, the positive predictive value (PPV) ranged from 29 to 100 percent, and the negative predictive value (NPV) ranged from 40 to 100 percent. The positive likelihood ratio (LR+) ranged from 0.98 to 3.00, and the negative likelihood ratio (LR-) ranged from 0.18 to 1.03. Using the summary sensitivity and specificity of 62 and 68 percent, respectively, we calculated an overall PPV of 57 percent and NPV of 72 percent. Similarly, we calculated summary LR+ of 1.94 and LR- of 0.56.

## Accuracy of ECG in 10 Good-Quality Studies

Next, we evaluated the accuracy of ECG compared with coronary angiography in the 10 good-quality studies. In these studies, sensitivity varied from 32 to 91 percent, and specificity varied from 46 to 81 percent; the median sensitivity was 71 percent, and the median specificity was 58 percent. Figure 6 presents forest plots of the individual study estimates of sensitivity and specificity of ECG in 10 good-quality studies for diagnosing CAD in women with no known CAD.

**Figure 6. Accuracy of ECG in 10 good-quality studies in women with no known CAD**

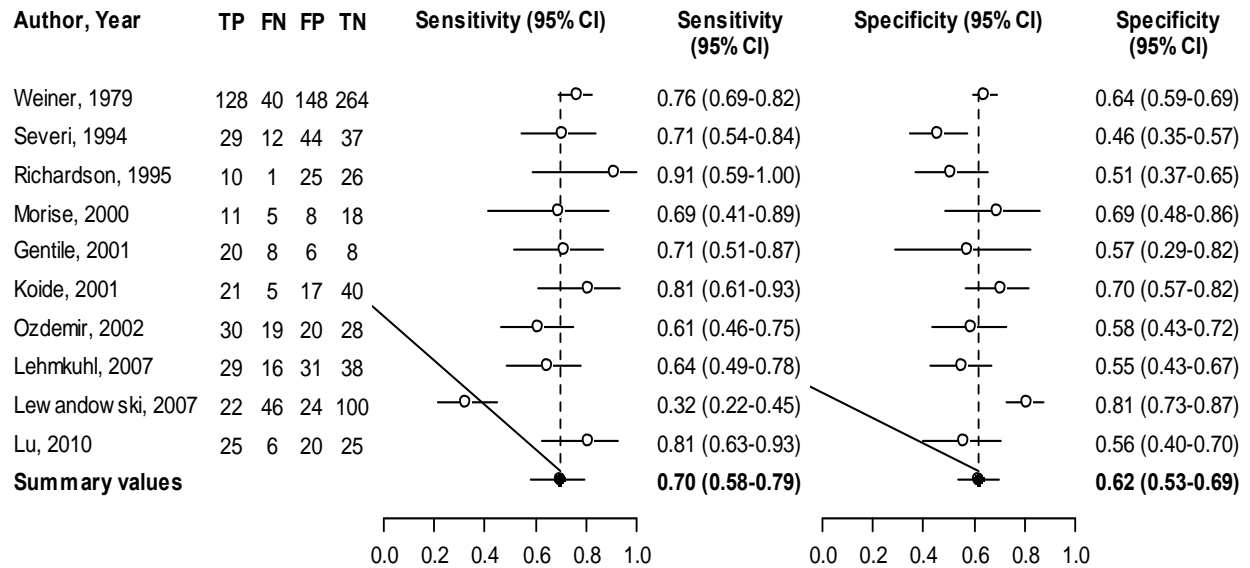
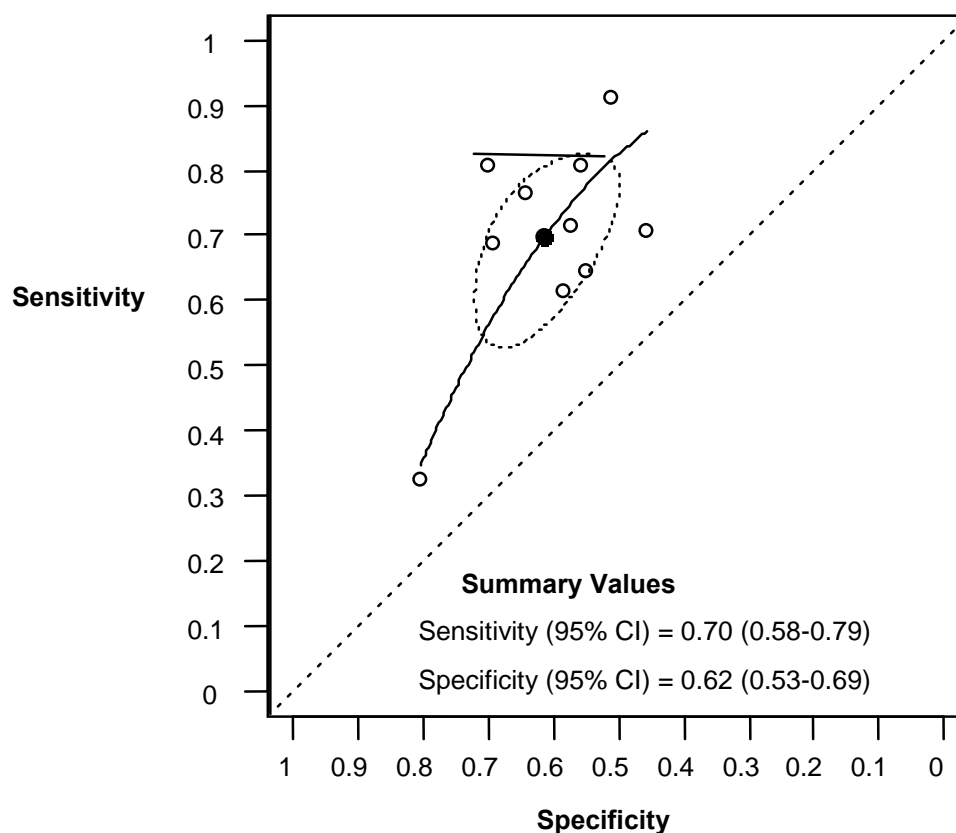


Figure 7 presents an SROC curve with an average sensitivity of 70 percent (95% CI, 58 to 79 percent) and specificity of 62 percent (95% CI, 53 to 69 percent).

**Figure 7. SROC curve for ECG in 10 good-quality studies in women with no known CAD**



The prevalence of CAD in these 10 good-quality studies ranged from 18 to 67 percent with a mean prevalence of 38 percent. In the individual studies, PPV ranged from 29 to 77 percent, and NPV ranged from 50 to 96 percent. LR+ ranged from 1.30 to 2.71 and LR- from 0.18 to 0.84. Using the summary sensitivity and specificity of 70 and 62 percent, respectively, we calculated an overall PPV of 53 percent and NPV of 77 percent. Similarly, we calculated summary LR+ of 1.84 and LR- of 0.48.

### **Secondary Analysis: Mixed Population of Women With Known and No Known CAD**

We performed a secondary analysis where we expanded our inclusion criteria to include studies whose patient population included a mix of women with known CAD and women with no known CAD. This expanded inclusion criteria allowed an additional 12 studies to be included in the analysis and an additional 83 patients from one study (totaling 41 studies). The 41 studies represent findings on ECG use in 4946 women (sample size ranging from 10 to 613 women). Of these 41 studies, 13 were good quality, 22 were fair quality, and 6 were poor quality (Table 2).

In these 41 studies, sensitivity varied from 26 to 96 percent, and specificity varied from 1 to 100 percent; the median sensitivity was 61 percent, and the median specificity was 65 percent. Figure 8 presents forest plots of the individual study estimates of sensitivity and specificity of ECG for diagnosing CAD in women from mixed populations.

**Figure 8. Accuracy of ECG in women from mixed populations**

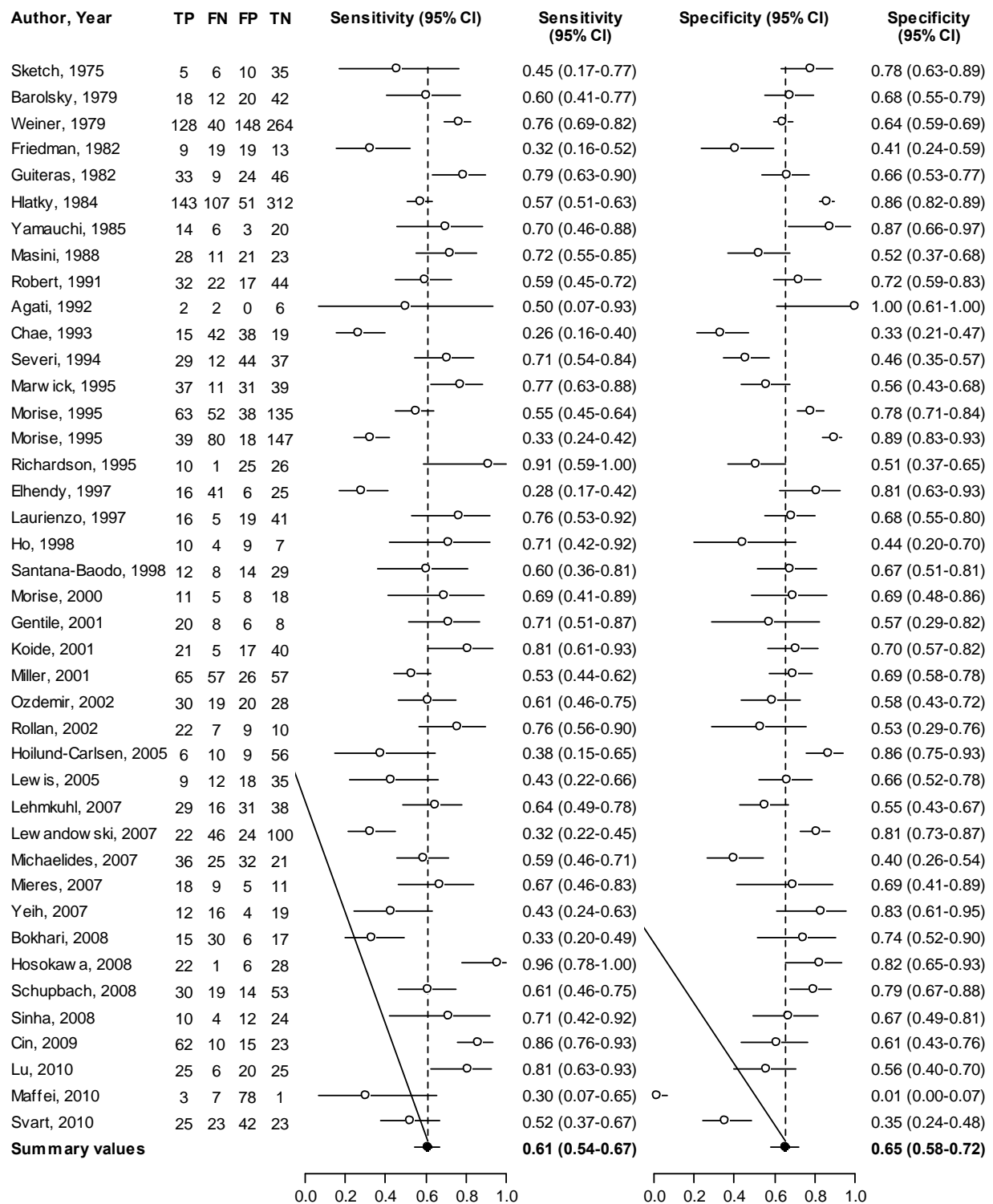
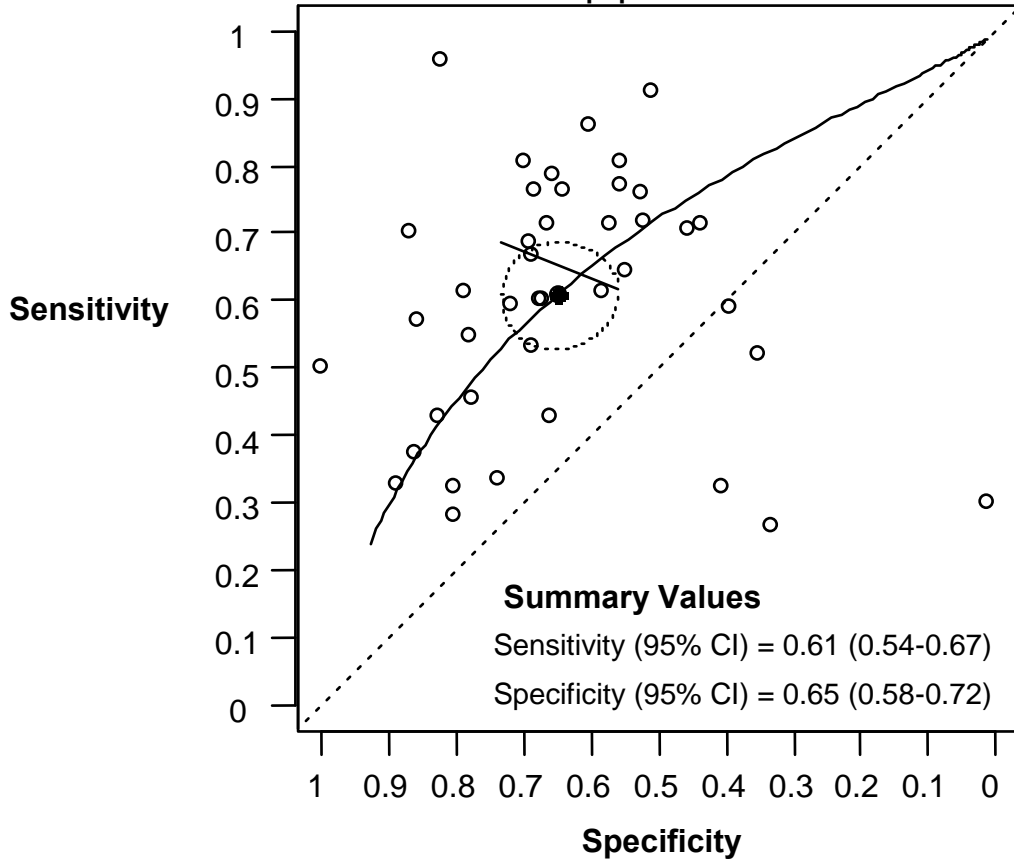


Figure 9 presents an SROC curve demonstrating an average sensitivity of 61 percent (95% CI, 54 to 67 percent) and specificity of 65 percent (95% CI, 58 to 72 percent).

**Figure 9. SROC curve for ECG in women from mixed populations**



The prevalence of CAD in these 41 studies ranged from 11 to 67 percent with a mean prevalence of 42 percent. In the individual studies, PPV ranged from 4 to 100 percent, and NPV ranged from 1 to 100 percent. LR+ ranged from 0.30 to 5.42 and LR- from 0.05 to 55.3. Using the summary sensitivity and specificity of 61 and 65 percent, respectively, we calculated an overall PPV of 56 percent and NPV of 70 percent. Similarly, we calculated summary LR+ of 1.74 and LR- of 0.60.

### **Accuracy of ECG in 13 Good-Quality Studies**

Next, we evaluated the accuracy of ECG compared with coronary angiography in the 13 good-quality studies. In these studies, sensitivity varied from 26 to 91 percent, and specificity varied from 33 to 81 percent; the median sensitivity was 71 percent, and the median specificity was 58 percent. Figure 10 presents forest plots of the individual study estimates of sensitivity and specificity of ECG in 13 good-quality studies for diagnosing CAD in women from mixed populations.



**Figure 10. Accuracy of ECG in 13 good-quality studies in women from mixed populations**

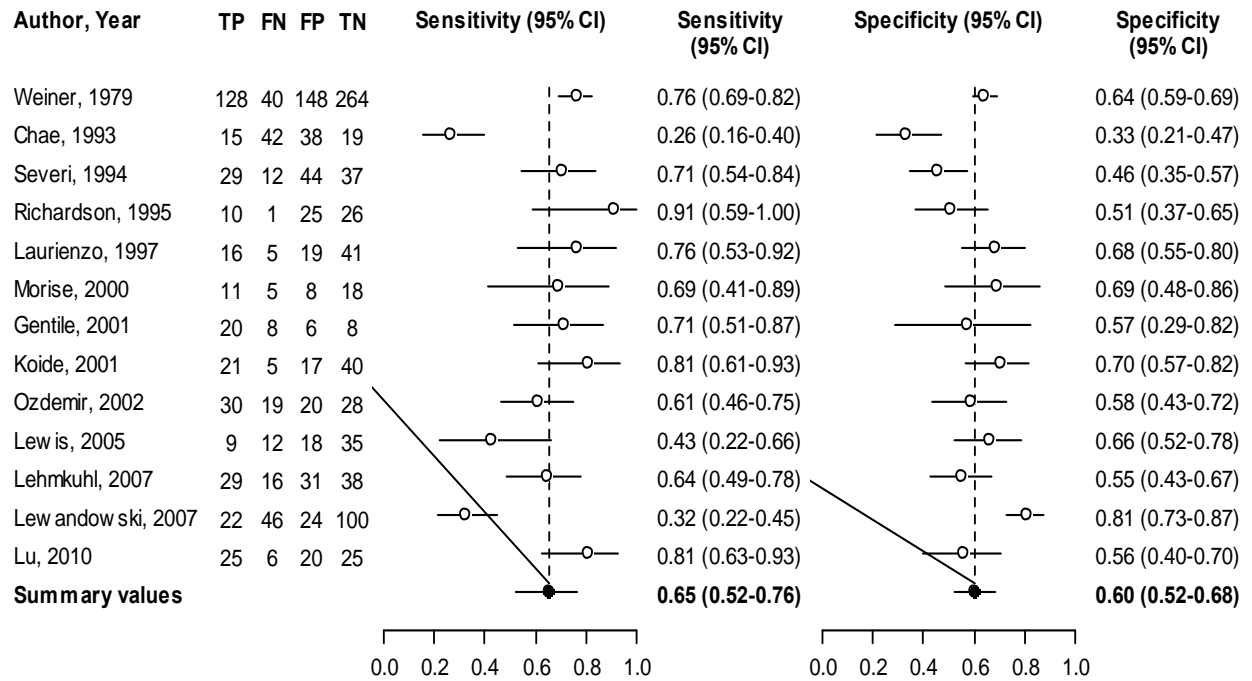
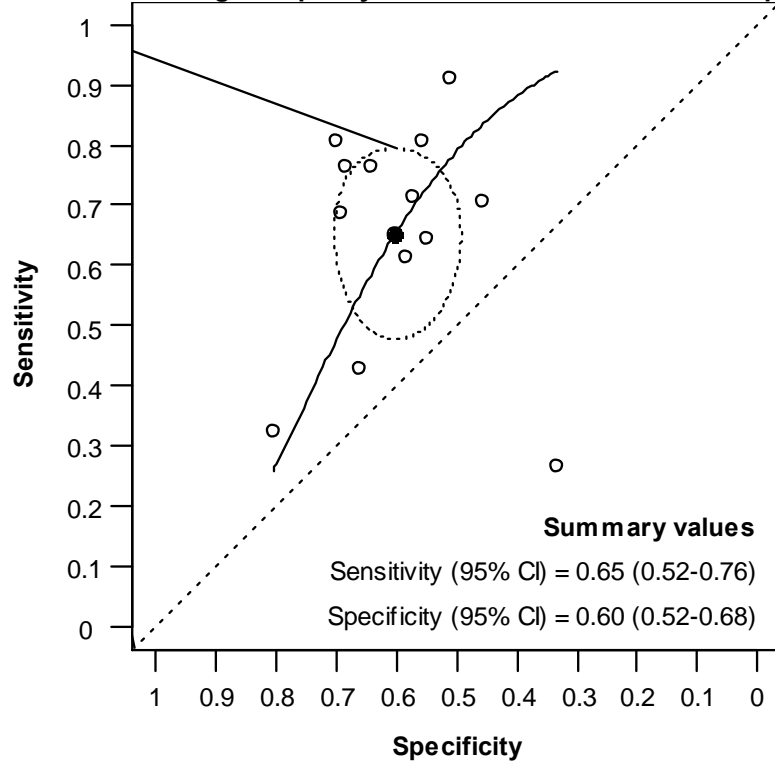


Figure 11 presents an SROC curve demonstrating an average sensitivity of 65 percent (95% CI, 52 to 76 percent) and specificity of 60 percent (95% CI, 52 to 68 percent).

**Figure 11. SROC curve for ECG in 13 good-quality studies in women from mixed populations**



The prevalence of CAD in these 13 good-quality studies ranged from 18 to 67 percent with a mean prevalence of 37 percent. In the individual studies, PPV ranged from 28 to 77 percent, and NPV ranged from 31 to 96 percent. LR+ ranged from 0.39 to 2.71 and LR- from 0.18 to 2.21. Using the summary sensitivity and specificity of 65 and 60 percent, respectively, we calculated an overall PPV of 49 percent and NPV of 75 percent. Similarly, we calculated summary LR+ of 1.62 and LR- of 0.58.

**Table 2. Summary of accuracy data evaluating ECG for diagnosing CAD**

| Study                                   | Patient Mix  | Quality | Patients (N) | Cath % | Prevalence % | Sensitivity % (95% CI) | Specificity % (95% CI) | PPV % (95% CI) | NPV % (95% CI) | LR+ (95% CI)     | LR- (95% CI)     |
|---|--------------|---------|--------------|--------|--------------|------------------------|------------------------|----------------|----------------|------------------|------------------|
| Lu, et al., 2010 <sup>28</sup>          | No known CAD | Good    | 76 (women)   | ≥ 50   | 41           | 81 (63-93)             | 56 (40-70)             | 56 (41-70)     | 81 (67-95)     | 1.81 (1.25-2.63) | 0.35 (0.16-0.75) |
| Lewandowski, et al., 2007 <sup>41</sup> | No known CAD | Good    | 192 (women)  | ≥ 50   | 35           | 32 (22-45)             | 81 (73-87)             | 48 (33-62)     | 68 (61-76)     | 1.67 (1.02-2.75) | 0.84 (0.70-1.01) |
|   |              |         | 359 (men)    |        | 75           | 20 (16-25)             | 93 (86-98)             | 90 (82-98)     | 28 (23-34)     | 3.06 (1.36-6.86) | 0.85 (0.79-0.93) |
| Ozdemir, et al., 2002 <sup>42</sup>     | No known CAD | Good    | 97 (women)   | ≥ 50   | 51           | 61 (46-75)             | 58 (43-72)             | 60 (46-74)     | 60 (46-74)     | 1.47 (0.98-2.20) | 0.66 (0.43-1.02) |
| Gentile, et al., 2001 <sup>43</sup>     | No known CAD | Good    | 42 (women)   | ≥ 60   | 67           | 71 (51-87)             | 57 (29-82)             | 77 (61-93)     | 50 (26-75)     | 1.67 (0.87-3.20) | 0.50 (0.24-1.05) |
|   |              |         | 90 (men)     |        | 89           | 90 (81-96)             | 60 (26-88)             | 95 (90-100)    | 43 (17-69)     | 2.25 (1.05-4.82) | 0.17 (0.07-0.38) |
| Koide, et al., 2001 <sup>44</sup>       | No known CAD | Good    | 83 (women)   | ≥ 50   | 31           | 81 (61-93)             | 70 (57-82)             | 55 (39-71)     | 89 (80-98)     | 2.71 (1.74-4.21) | 0.27 (0.12-0.61) |
|   |              |         | 190 (men)    |        | 53           | 62 (52-72)             | 74 (64-83)             | 73 (64-83)     | 63 (54-73)     | 2.41 (1.65-3.54) | 0.51 (0.38-0.67) |
| Morise, et al., 2000 <sup>45</sup>      | No known CAD | Good    | 42 (women)   | ≥ 50   | 38           | 69 (41-89)             | 69 (48-86)             | 58 (36-80)     | 78 (61-95)     | 2.23 (1.15-4.34) | 0.45 (0.21-0.98) |
| Lehmkuhl, et al., 2007 <sup>27</sup>    | No known CAD | Good    | 114 (women)  | ≥ 50   | 39           | 64 (49-78)             | 55 (43-67)             | 48 (36-61)     | 70 (58-83)     | 1.43 (1.02-2.01) | 0.65 (0.41-1.01) |
| Severi, et al., 1994 <sup>46</sup>      | No known CAD | Good    | 122 (women)  | ≥ 75   | 34           | 71 (54-84)             | 46 (35-57)             | 40 (29-51)     | 76 (63-88)     | 1.30 (0.98-1.72) | 0.64 (0.38-1.09) |
|   |              |         | 307 (men)    |        | 67           | 75 (69-81)             | 56 (46-66)             | 77 (72-83)     | 53 (43-62)     | 1.70 (1.35-2.15) | 0.45 (0.33-0.60) |

**Table 2. Summary of accuracy data evaluating ECG for diagnosing CAD (continued)**

| Study                                       | Patient Mix  | Quality | Patients (N) | Cath % | Prevalence % | Sensitivity % (95% CI) | Specificity % (95% CI) | PPV % (95% CI) | NPV % (95% CI)    | LR+ (95% CI)     | LR- (95% CI)     |
|---|--------------|---------|--------------|--------|--------------|------------------------|------------------------|----------------|-------------------|------------------|------------------|
| Richards, et al., 1995 <sup>47</sup>        | No known CAD | Good    | 62 (women)   | ≥ 50   | 18           | 91 (59-100)            | 51 (37-65)             | 29 (14-44)     | 96 (89-100)       | 1.85 (1.32-2.60) | 0.18 (0.03-1.18) |
| Weiner, et al., 1979 <sup>48</sup>          | No known CAD | Good    | 580 (women)  | ≥ 70   | 29           | 76 (69-82)             | 64 (59-69)             | 46 (40-52)     | 87 (83-91)        | 2.12 (1.82-2.47) | 0.37 (0.28-0.49) |
|   |              |         | 1465 (men)   | 70     | 80 (77-82)   | 74 (70-78)             | 88 (86-90)             | 61 (57-65)     | 3.08 (2.62-3.62)  | 0.27 (0.24-0.31) |                  |
| Schupbach, et al., 2008 <sup>49a</sup>      | No known CAD | Fair    | 116 (women)  | ≥ 50   | 42           | 61 (46-75)             | 79 (67-88)             | 68 (54-82)     | 74 (63-84)        | 2.93 (1.75-4.91) | 0.49 (0.34-0.71) |
|   |              |         | 216 (men)    | 73     | 74 (67-80)   | 86 (78-92)             | 92 (87-97)             | 47 (38-57)     | 4.20 (2.28-7.74)  | 0.41 (0.32-0.52) |                  |
| Yeih, et al., 2007 <sup>50</sup>            | No known CAD | Fair    | 51 (women)   | ≥ 50   | 55           | 43 (24-63)             | 83 (61-95)             | 75 (54-96)     | 54 (38-71)        | 2.46 (0.92-6.62) | 0.69 (0.48-1.00) |
| Michaelides, et al., 2007 <sup>51</sup>     | No known CAD | Fair    | 114 (women)  | ≥ 50   | 54           | 59 (46-71)             | 40 (26-54)             | 53 (41-65)     | 46 (31-60)        | 0.98 (0.72-1.32) | 1.03 (0.66-1.62) |
| Mieres, et al., 2007 <sup>52</sup>          | No known CAD | Fair    | 42 (women)   | ≥ 50   | 64           | 67 (46-83)             | 69 (41-89)             | 78 (61-95)     | 55 (33-77)        | 2.13 (0.98-4.63) | 0.48 (0.26-0.91) |
| Hoilund-Carlsen, et al., 2005 <sup>53</sup> | No known CAD | Fair    | 81 (women)   | ≥ 50   | 20           | 38 (15-65)             | 86 (75-93)             | 40 (15-65)     | 85 (76-93)        | 2.71 (1.13-6.51) | 0.73 (0.49-1.07) |
|   |              |         | 105 (men)    | 44     | 74 (59-86)   | 92 (81-97)             | 87 (77-98)             | 82 (73-91)     | 8.72 (3.71-20.53) | 0.29 (0.17-0.47) |                  |
| Rollan, et al., 2002 <sup>54</sup>          | No known CAD | Fair    | 48 (women)   | ≥ 50   | 60           | 76 (56-90)             | 53 (29-76)             | 71 (55-87)     | 59 (35-82)        | 1.60 (0.96-2.68) | 0.46 (0.21-0.99) |
| Miller, et al., 2001 <sup>29</sup>          | No known CAD | Fair    | 205 (women)  | ≥ 50   | 60           | 53 (44-62)             | 69 (58-78)             | 71 (62-81)     | 50 (41-59)        | 1.70 (1.19-2.44) | 0.68 (0.54-0.86) |
|   |              |         | 838 (men)    | 75     | 63 (59-67)   | 74 (68-80)             | 88 (85-91)             | 40 (35-45)     | 2.44 (1.92-3.09)  | 0.50 (0.44-0.57) |                  |
| Marwick, et al., 1995 <sup>55</sup>         | No known CAD | Fair    | 118 (women)  | ≥ 50   | 41           | 77 (63-88)             | 56 (43-68)             | 54 (43-66)     | 78 (67-89)        | 1.74 (1.28-2.36) | 0.41 (0.24-0.72) |

**Table 2. Summary of accuracy data evaluating ECG for diagnosing CAD (continued)**

| Study                                     | Patient Mix  | Quality | Patients (N) | Cath % | Prevalence % | Sensitivity % (95% CI) | Specificity % (95% CI) | PPV % (95% CI) | NPV % (95% CI) | LR+ (95% CI)      | LR- (95% CI)     |
|---|--------------|---------|--------------|--------|--------------|------------------------|------------------------|----------------|----------------|-------------------|------------------|
| Agati, et al., 1992 <sup>56</sup>         | No known CAD | Fair    | 10 (women)   | ≥ 70   | 40           | 50 (7-93)              | 100 (61-100)           | 100 (NA-100)   | 75 (45-100)    | NA                | 0.50 (0.19-1.33) |
|   |              |         | 22 (men)     |        | 82           | 72 (47-90)             | 75 (19-99)             | 93 (79-100)    | 38 (4-71)      | 2.89 (0.52-16.16) | 0.37 (0.15-0.94) |
| Barolsky, et al., 1979 <sup>57</sup>      | No known CAD | Fair    | 92 (women)   | ≥ 75   | 33           | 60 (41-77)             | 68 (55-79)             | 47 (31-63)     | 78 (67-89)     | 1.86 (1.17-2.96)  | 0.59 (0.37-0.95) |
|   |              |         | 85 (men)     |        | 36           | 65 (45-81)             | 89 (77-96)             | 77 (61-93)     | 81 (71-91)     | 5.81 (2.61-12.90) | 0.40 (0.25-0.65) |
| Morise, et al., 1995 <sup>58</sup>        | No known CAD | Fair    | 284 (women)  | ≥ 50   | 42           | 33 (24-42)             | 89 (83-93)             | 68 (56-80)     | 65 (59-71)     | 3.00 (1.81-4.98)  | 0.75 (0.66-0.86) |
|   |              |         | 504 (men)    |        | 63           | 40 (35-46)             | 40 (33-47)             | 53 (47-59)     | 28 (23-33)     | 0.66 (0.55-0.79)  | 1.51 (1.24-1.84) |
| Robert, et al., 1991 <sup>59</sup>        | No known CAD | Fair    | 115 (women)  | ≥ 50   | 47           | 59 (45-72)             | 72 (59-83)             | 65 (52-79)     | 67 (55-78)     | 2.12 (1.34-3.37)  | 0.56 (0.40-0.81) |
| Guiteras, et al., 1982 <sup>60</sup>      | No known CAD | Fair    | 112 (women)  | ≥ 70   | 38           | 79 (63-90)             | 66 (53-77)             | 58 (45-71)     | 84 (74-93)     | 2.29 (1.60-3.29)  | 0.33 (0.18-0.60) |
| Masini, et al., 1988 <sup>61</sup>        | No known CAD | Fair    | 68 (women)   | ≥ 70   | 38           | 69 (48-86)             | 52 (36-68)             | 47 (31-63)     | 73 (58-89)     | 1.45 (0.97-2.19)  | 0.59 (0.31-1.12) |
| Santana-Baodo, et al., 1998 <sup>68</sup> | No known CAD | Fair    | 63 (women)   | ≥ 50   | 32           | 60 (36-81)             | 67 (51-81)             | 46 (27-65)     | 78 (65-92)     | 1.84 (1.05-3.22)  | 0.59 (0.33-1.05) |
|   |              |         | 100 (men)    |        | 80           | 69 (57-79)             | 80 (56-94)             | 93 (87-100)    | 39 (24-54)     | 3.43 (1.41-8.36)  | 0.39 (0.26-0.58) |
| Bokhari, et al., 2008 <sup>62</sup>       | No known CAD | Poor    | 68 (women)   | ≥ 50   | 66           | 33 (20-49)             | 74 (52-90)             | 71 (52-91)     | 36 (22-50)     | 1.28 (0.57-2.85)  | 0.90 (0.66-1.24) |
|   |              |         | 150 (men)    |        | 65           | 37 (27-47)             | 94 (84-99)             | 92 (84-100)    | 44 (35-53)     | 6.37 (2.06-19.69) | 0.67 (0.57-0.79) |
| Sinha, et al., 2008 <sup>63</sup>         | No known CAD | Poor    | 50 (women)   | ≥ 50   | 28           | 71 (42-92)             | 67 (49-81)             | 45 (25-66)     | 86 (73-99)     | 2.14 (1.21-3.78)  | 0.43 (0.18-1.01) |

**Table 2. Summary of accuracy data evaluating ECG for diagnosing CAD (continued)**

| Study                                 | Patient Mix  | Quality | Patients (N) | Cath % | Prevalence % | Sensitivity % (95% CI) | Specificity % (95% CI) | PPV % (95% CI) | NPV % (95% CI)    | LR+ (95% CI)      | LR- (95% CI)      |
|---------------------------------------|--------------|---------|--------------|--------|--------------|------------------------|------------------------|----------------|-------------------|-------------------|-------------------|
| Sketch, et al., 1975 <sup>64</sup>    | No known CAD | Poor    | 56 (women)   | ≥ 50   | 20           | 45 (17-77)             | 78 (63-89)             | 33 (9-57)      | 85 (75-96)        | 2.05 (0.88-4.77)  | 0.70 (0.40-1.23)  |
|                                       |              |         | 195 (men)    | 57     | 51 (42-61)   | 94 (87-98)             | 92 (85-99)             | 59 (51-68)     | 8.63 (3.62-20.57) | 0.52 (0.42-0.63)  |                   |
| Morise, et al., 1995 <sup>65</sup>    | No known CAD | Poor    | 288 (women)  | ≥ 50   | 40           | 55 (45-64)             | 78 (71-84)             | 62 (53-72)     | 72 (66-79)        | 2.50 (1.80-3.46)  | 0.58 (0.47-0.72)  |
|                                       |              |         | 577 (men)    | 66     | 58 (53-63)   | 79 (73-85)             | 84 (80-89)             | 49 (44-55)     | 2.77 (2.08-3.69)  | 0.53 (0.46-0.61)  |                   |
| Lewis, et al., 2005 <sup>38</sup>     | Mixed        | Good    | 74 (women)   | ≥ 50   | 28           | 43 (22-66)             | 66 (52-78)             | 33 (16-51)     | 74 (62-87)        | 1.26 (0.68-2.35)  | 0.87 (0.57-1.31)  |
| Laurienzo, et al., 1997 <sup>39</sup> | Mixed        | Good    | 81 (women)   | ≥ 70   | 26           | 76 (53-92)             | 68 (55-80)             | 46 (29-62)     | 89 (80-98)        | 2.41 (1.55-3.74)  | 0.35 (0.16-0.76)  |
| Chae, et al., 1993 <sup>40</sup>      | Mixed        | Good    | 114 (women)  | ≥ 50   | 50           | 26 (16-40)             | 33 (21-47)             | 28 (16-40)     | 31 (20-43)        | 0.40 (0.25-0.63)  | 2.21 (1.48-3.29)  |
| Hosokawa, et al., 2008 <sup>33a</sup> | Mixed        | Fair    | 57 (women)   | ≥ 70   | 40           | 96 (78-100)            | 82 (65-93)             | 79 (63-94)     | 97 (90-100)       | 5.42 (2.61-11.26) | 0.053 (0.01-0.36) |
|                                       |              |         | 132 (men)    | 41     | 94 (85-99)   | 88 (79-95)             | 85 (76-94)             | 96 (91-100)    | 8.19 (4.41-15.18) | 0.06 (0.02-0.19)  |                   |
| Svart, et al., 2010 <sup>34</sup>     | Mixed        | Fair    | 113 (women)  | ≥ 50   | 42           | 52 (37-67)             | 35 (24-48)             | 37 (26-49)     | 50 (36-64)        | 0.81 (0.58-1.12)  | 1.35 (0.87-2.11)  |
| Elhendy, et al., 1997 <sup>35</sup>   | Mixed        | Fair    | 88 (women)   | ≥ 50   | 65           | 28 (17-42)             | 81 (63-93)             | 73 (54-91)     | 38 (26-50)        | 1.45 (0.63-3.33)  | 0.89 (0.70-1.13)  |
|                                       |              |         | 177 (men))   | 81     | 40 (32-49)   | 85 (68-95)             | 92 (85-99)             | 25 (17-32)     | 2.66 (1.16-6.11)  | 0.70 (0.58-0.86)  |                   |
| Yamauchi, et al., 1985 <sup>36</sup>  | Mixed        | Fair    | 43 (women)   | ≥ 75   | 47           | 70 (46-88)             | 87 (66-97)             | 82 (64-100)    | 77 (61-93)        | 5.37 (1.80-16.02) | 0.35 (0.17-0.69)  |
|                                       |              |         | 90 (men)     | 71     | 70 (58-81)   | 92 (75-99)             | 96 (90-100)            | 56 (41-71)     | 9.14 (2.39-34.94) | 0.32 (0.22-0.48)  |                   |
| Masini, et al., 1988 <sup>61</sup>    | Mixed        | Fair    | 83 (women)   | ≥ 70   | 47           | 72 (55-85)             | 52 (37-68)             | 57 (43-71)     | 68 (52-83)        | 1.50 (1.04-2.17)  | 0.54 (0.30-0.96)  |
| Friedman, et al., 1982 <sup>37</sup>  | Mixed        | Fair    | 60 (women)   | ≥ 70   | 47           | 32 (16-52)             | 41 (24-59)             | 32 (15-49)     | 41 (24-58)        | 0.54 (0.29-1.00)  | 1.67 (1.02-2.73)  |

**Table 2. Summary of accuracy data evaluating ECG for diagnosing CAD (continued)**

| Study                              | Patient Mix | Quality | Patients (N) | Cath % | Prevalence % | Sensitivity % (95% CI) | Specificity % (95% CI) | PPV % (95% CI) | NPV % (95% CI) | LR+ (95% CI)     | LR- (95% CI)     |
|------------------------------------|-------------|---------|--------------|--------|--------------|------------------------|------------------------|----------------|----------------|------------------|------------------|
| Maffei, et al., 2010 <sup>66</sup> | Mixed       | Fair    | 89 (women)   | ≥ 50   | 11           | 30 (7-65)              | 1 (0-7)                | 4 (0-8)        | 13 (0-35)      | 0.30 (0.12-0.78) | 55.3 (7.56-404)  |
|                                    |             |         | 88 (men)     |        |              | 50 (25-75)             | 35 (24-47)             | 15 (5-24)      | 76 (61-90)     | 0.76 (0.46-1.28) | 1.44 (0.80-2.58) |
| Ho, et al., 1998 <sup>67</sup>     | Mixed       | Fair    | 30 (women)   | ≥ 50   | 47           | 71 (42-92)             | 44 (20-70)             | 53 (30-75)     | 64 (35-92)     | 1.27 (0.74-2.19) | 0.65 (0.24-1.77) |
| Cin, et al., 2000 <sup>31</sup>    | Mixed       | Poor    | 110 (women)  | ≥ 50   | 65           | 86 (76-93)             | 61 (43-76)             | 81 (72-89)     | 70 (54-85)     | 2.18 (1.46-3.27) | 0.23 (0.12-0.43) |
| Hlatky, et al., 1984 <sup>32</sup> | Mixed       | Poor    | 613 (women)  | NR     | 41           | 57 (51-63)             | 86 (82-89)             | 74 (68-80)     | 74 (70-79)     | 4.07 (3.09-5.37) | 0.50 (0.43-0.58) |
|                                    |             |         | 1656 (men)   |        |              | 72 (70-75)             | 83 (79-86)             | 92 (90-94)     | 53 (49-57)     | 4.28 (3.47-5.27) | 0.33 (0.30-0.37) |

<sup>a</sup>Resting ECG

Abbreviations: CAD = coronary artery disease; Cath % = % stenosis defined to be positive for CAD on diagnostic cardiac catheterization (coronary angiography); CI = confidence interval; LR+ = positive likelihood ratio; LR- = negative likelihood ratio; NPV = negative predictive value; NR = not reported; PPV = positive predictive value

## ECHO

We identified 22 studies evaluating the accuracy of exercise/stress ECHO compared with coronary angiography (Table 3).<sup>27,28,35,39,41,46,54-56,61,67,69-79</sup> Fourteen of these studies reported accuracy data in women with no known CAD, and these are the studies used in our primary analyses. In our secondary analyses, we evaluated the accuracy of the ECHO in diagnosing CAD including an additional 8 studies that reported findings with mixed populations of known and no known CAD as well as additional patients from one study<sup>61</sup> that had data for those with no known CAD. None of the identified ECHO studies used contrast.

### Primary Analysis: Population of Women With No Known CAD

The 14 studies represent findings on ECHO use in 1289 women (sample size ranging from 14 to 192 women). Of these studies, five were good-quality, eight were fair-quality, and one was poor-quality. Sensitivity varied from 57 to 90 percent, specificity varied from 37 to 96 percent; the median sensitivity was 79 percent, and the median specificity was 82 percent. Figure 12 presents forest plots of the individual study estimates of sensitivity and specificity of ECHO for diagnosing CAD in women with no known CAD.

**Figure 12. Accuracy of ECHO in women with no known CAD**

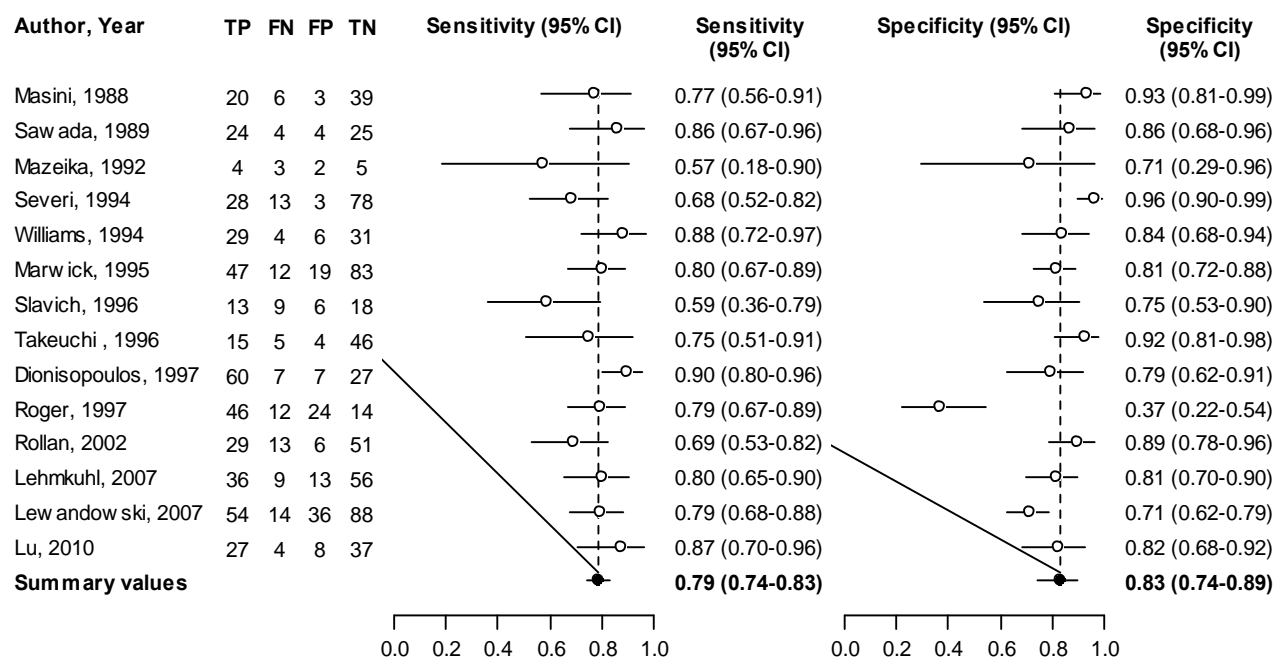
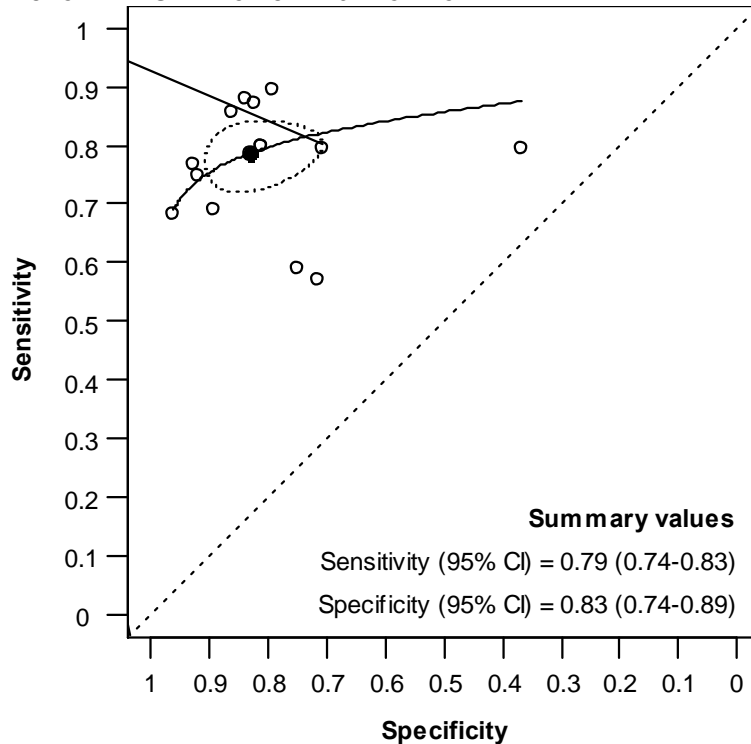




Figure 13 presents an SROC curve with an average sensitivity of 79 percent (95% CI, 74 to 83 percent) and specificity of 83 (95% CI, 74 to 89 percent).

**Figure 13. SROC curve for ECHO in women with no known CAD**



The prevalence of CAD in these 14 studies ranged from 29 to 66 percent with a mean prevalence of 44 percent. In the individual studies, PPV ranged from 60 to 90 percent, and NPV ranged from 37 to 96 percent. LR+ ranged from 1.25 to 18.44 and LR- from 0.13 to 0.60. Using the summary sensitivity and specificity of 79 and 83 percent, respectively, we calculated an overall PPV of 78 percent and NPV of 83 percent. Similarly, we calculated summary LR+ of 4.65 and LR- of 0.25.

### **Accuracy of ECHO in Five Good-Quality Studies**

Next, we evaluated the accuracy of ECHO compared with coronary angiography in the five good-quality studies. In these studies, sensitivity varied from 68 to 87 percent, specificity varied from 71 to 96 percent; the median sensitivity was 80 percent, and the median specificity was 82 percent. Figure 14 presents forest plots of the individual study estimates of sensitivity and specificity of ECHO for diagnosing CAD in women with no known CAD.

**Figure 14. Accuracy of ECHO in five good-quality studies in women with no known CAD**

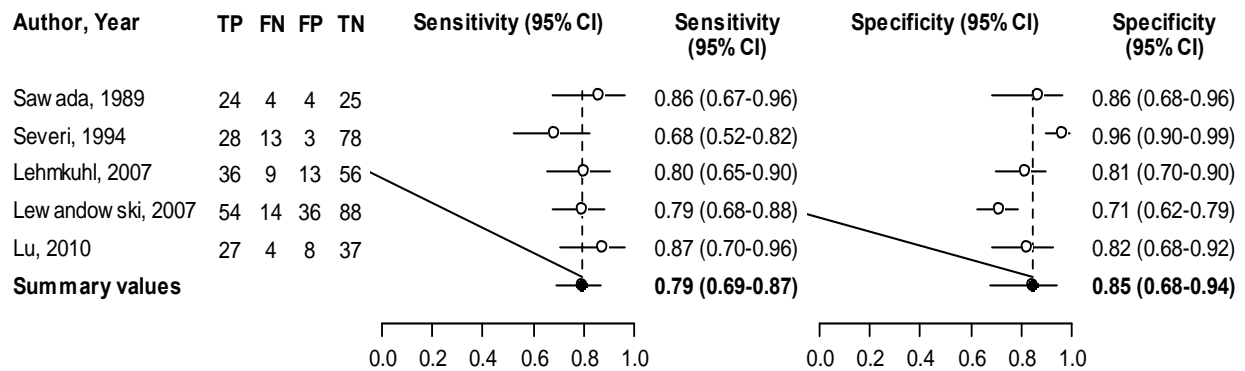
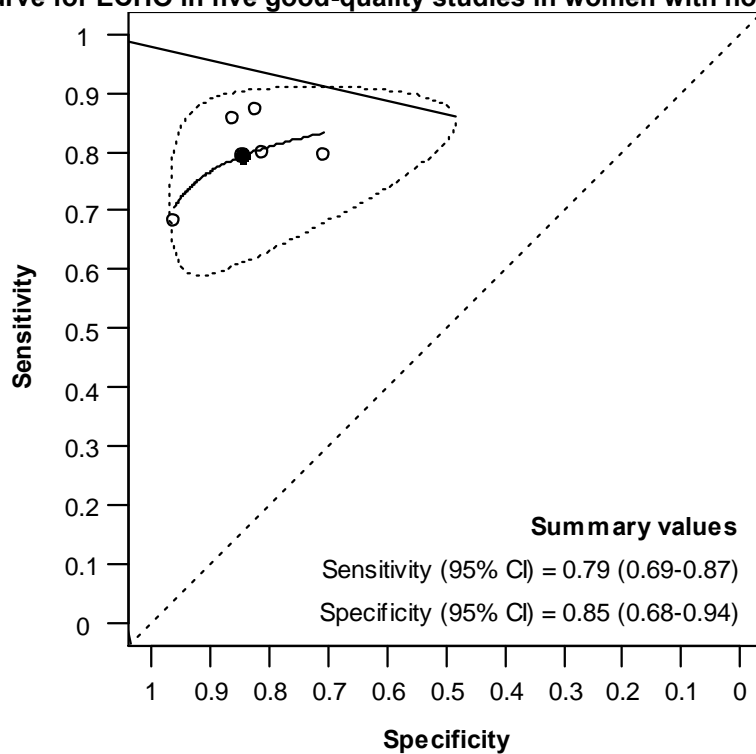


Figure 15 presents an SROC curve with an average sensitivity of 79 percent (95% CI, 69 to 87 percent) and specificity of 85 percent (95% CI, 68 to 94 percent).

**Figure 15. SROC curve for ECHO in five good-quality studies in women with no known CAD**



The prevalence of CAD in these five good-quality studies ranged from 34 to 49 percent with a mean prevalence of 40 percent. In the individual studies, PPV ranged from 60 to 90 percent, and NPV ranged from 86 to 90 percent. LR+ ranged from 2.73 to 18.44 and LR- from 0.16 to 0.33. Using the summary sensitivity and specificity of 79 and 85 percent, respectively, we calculated an overall PPV of 78 percent and NPV of 86 percent. Similarly, we calculated summary LR+ of 5.27 and LR- of 0.25.

## Secondary Analysis: Mixed Population of Women With Known and No Known CAD

We performed a secondary analysis where we expanded our inclusion criteria to include studies whose patient population included a mix of women with known CAD and women with no known CAD. This expanded inclusion criteria allowed an additional eight studies to be included in the analysis and an additional group of patients from one study (totaling 22 studies). The 22 studies represent findings on ECHO use in 1944 women (sample size ranging from 7 to 192 women). Of these 22 studies, 8 were good quality, 13 were fair quality, and 1 was poor quality (Table 3).

In these 22 studies, sensitivity varied from 40 to 93 percent, and specificity varied from 37 to 100 percent; the median sensitivity was 79 percent, and the median specificity was 84 percent. Figure 16 presents forest plots of the individual study estimates of sensitivity and specificity of ECHO for diagnosing CAD in women from mixed populations.

**Figure 16. Accuracy of ECHO in women from mixed populations**

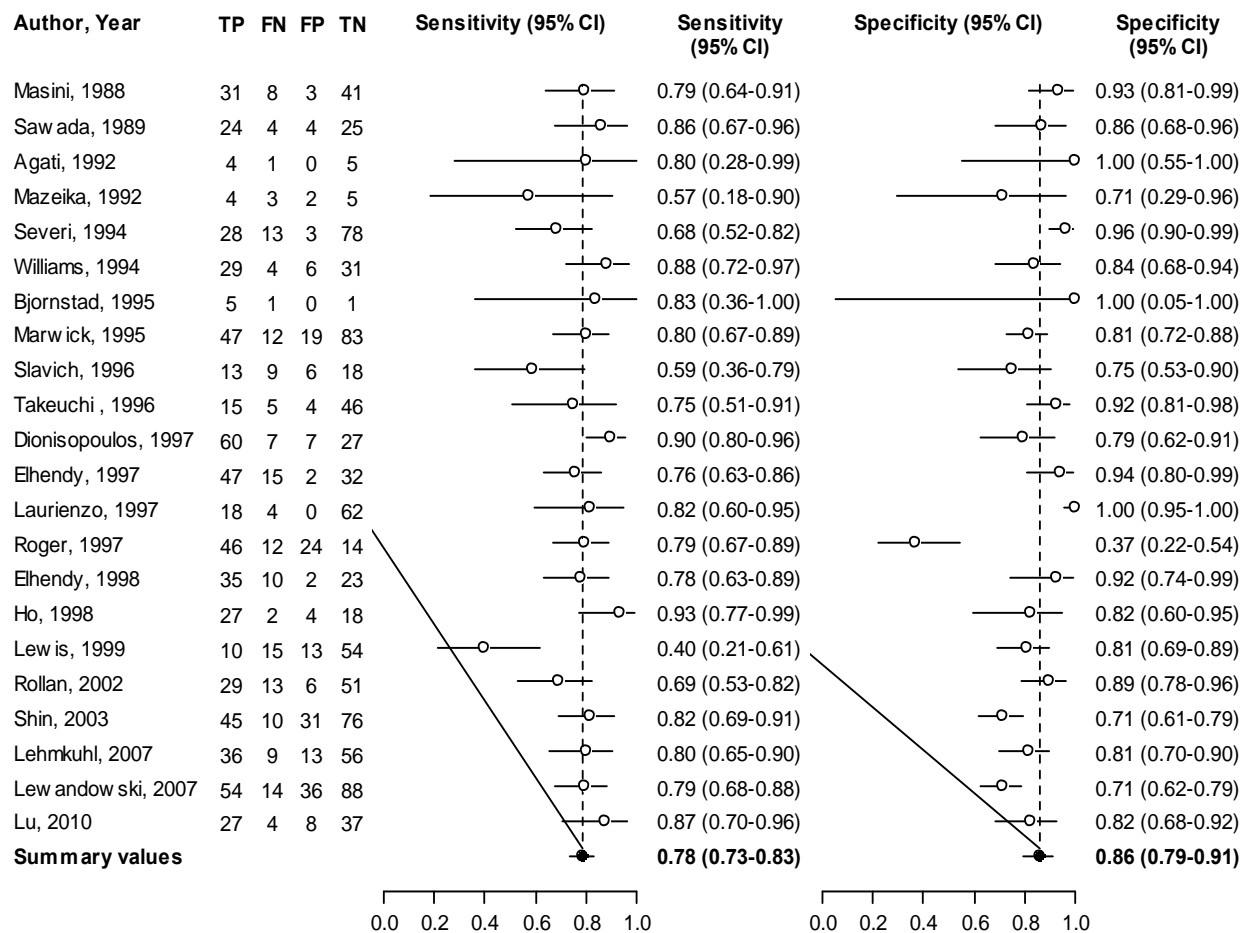
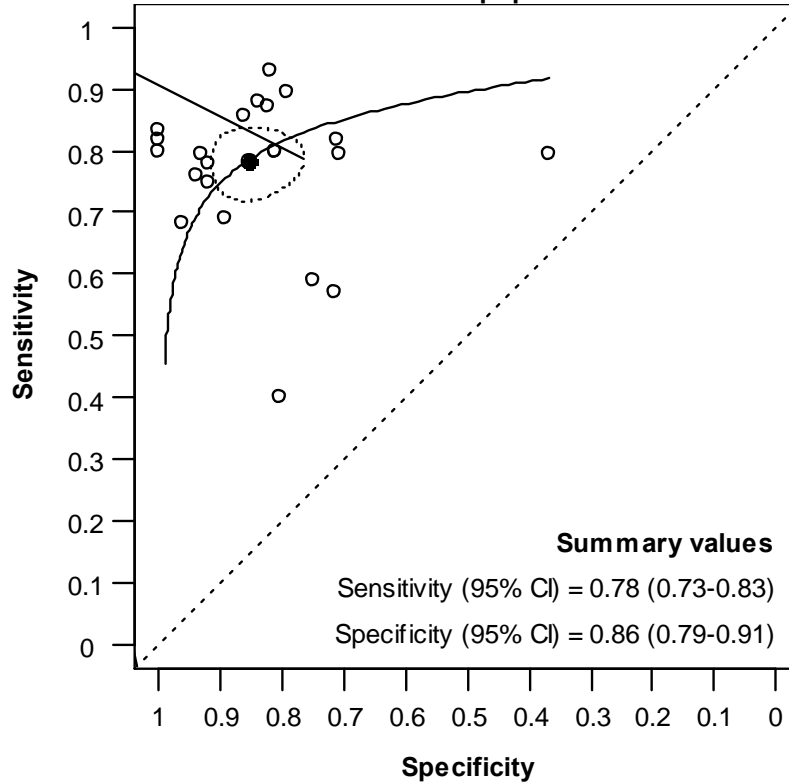


Figure 17 presents an SROC curve demonstrating an average sensitivity of 78 percent (95% CI, 73 to 83 percent) and specificity of 86 percent (95% CI, 79 to 91 percent).

**Figure 17. SROC curve for ECHO in women from mixed populations**



The prevalence of CAD in these 22 studies ranged from 26 to 86 percent with a mean prevalence of 46 percent. In the individual studies, PPV ranged from 43 to 100 percent, and NPV ranged from 37 to 100 percent. LR+ ranged from 1.25 to 18.44 and LR- from 0.08 to 0.74. Using the summary sensitivity and specificity of 78 and 86 percent, respectively, we calculated an overall PPV of 82 percent and NPV of 82 percent. Similarly, we calculated summary LR+ of 5.57 and LR- of 0.26.

### **Accuracy of ECHO in Eight Good-Quality Studies**

Next, we evaluated the accuracy of ECHO compared with coronary angiography in the eight good-quality studies. In these studies, sensitivity varied from 40 to 87 percent, and specificity varied from 71 to 100 percent; the median sensitivity was 80 percent, and the median specificity was 84 percent. Figure 18 presents forest plots of the individual study estimates of sensitivity and specificity of ECHO for diagnosing CAD in women from mixed populations.

**Figure 18. Accuracy of ECHO in eight good-quality studies in women from mixed populations**

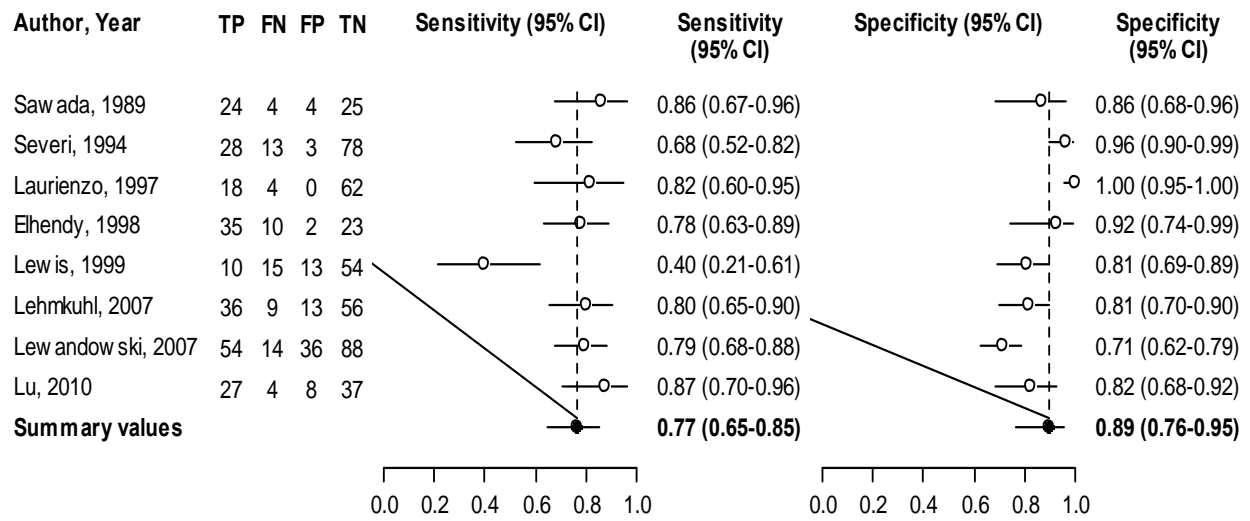
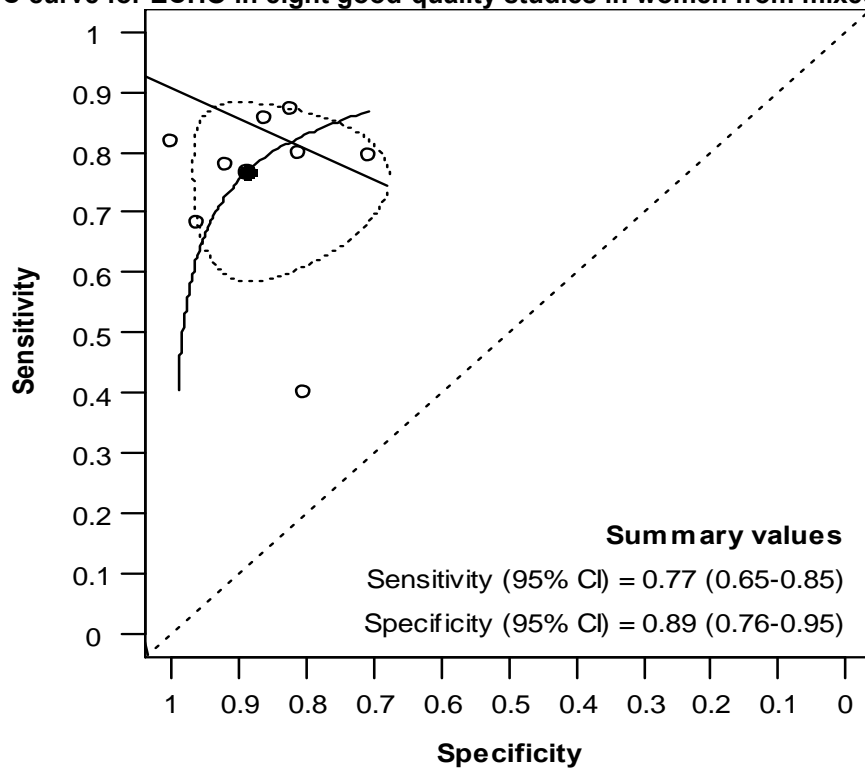


Figure 19 presents an SROC curve demonstrating an average sensitivity of 77 percent (95% CI, 65 to 85 percent) and specificity of 89 percent (95% CI, 76 to 95 percent).

**Figure 19. SROC curve for ECHO in eight good-quality studies in women from mixed populations**



The prevalence of CAD in these 8 good-quality studies ranged from 26 to 64 percent with a mean prevalence of 40 percent. In the individual studies, PPV ranged from 43 to 100 percent, and NPV ranged from 70 to 94 percent. LR+ ranged from 2.06 to 18.44 and LR- from 0.16 to 0.74. Using the summary sensitivity and specificity of 77 and 89 percent, respectively, we calculated an overall PPV of 82 percent and NPV of 85 percent. Similarly, we calculated summary LR+ of 7.0 and LR- of 0.26.

**Table 3. Summary of accuracy data evaluating ECHO for diagnosing CAD**

| Study                                     | Patient Mix  | Quality | Patients (N) | Cath % | Prevalence % | Sensitivity % (95% CI) | Specificity % (95% CI) | PPV % (95% CI) | NPV % (95% CI) | LR+ (95% CI)       | LR- (95% CI)     |
|---|--------------|---------|--------------|--------|--------------|------------------------|------------------------|----------------|----------------|--------------------|------------------|
| Lu, et al., 2010 <sup>28</sup>            | No known CAD | Good    | 76 (women)   | ≥ 50   | 41           | 87 (75-99)             | 82 (71-93)             | 77 (63-91)     | 90 (81-99)     | 4.90 (2.58-9.32)   | 0.16 (0.06-0.40) |
| Lewandowski, et al., 2007 <sup>41</sup>   | No known CAD | Good    | 192 (women)  | ≥ 50   | 35           | 79 (70-89)             | 71 (63-79)             | 60 (50-70)     | 86 (80-93)     | 2.74 (2.03-3.70)   | 0.29 (0.18-0.47) |
|   |              |         | 359 (men)    |        | 75           | 87 (82-91)             | 66 (55-76)             | 88 (84-92)     | 63 (53-73)     | 2.55 (1.91-3.41)   | 0.20 (0.14-0.28) |
| Lehmkuhl, et al., 2007 <sup>27</sup>      | No known CAD | Good    | 114 (women)  | ≥ 50   | 39           | 80 (68-92)             | 81 (72-90)             | 73 (61-86)     | 86 (78-95)     | 4.25 (2.55-7.08)   | 0.25 (0.14-0.45) |
| Severi, et al., 1994 <sup>46</sup>        | No known CAD | Good    | 122 (women)  | ≥ 75   | 34           | 68 (54-83)             | 96 (92-100)            | 90 (80-100)    | 86 (79-93)     | 18.44 (5.96-57.07) | 0.33 (0.21-0.52) |
|   |              |         | 307 (men)    |        | 67           | 76 (70-82)             | 84 (76-91)             | 91 (86-95)     | 64 (56-72)     | 4.85 (3.07-7.66)   | 0.28 (0.22-0.37) |
| Sawada, et al., 1989 <sup>73</sup>        | No known CAD | Good    | 57 (women)   | ≥ 50   | 49           | 86 (73-99)             | 86 (74-99)             | 86 (73-99)     | 86 (74-99)     | 6.21 (2.47-15.63)  | 0.17 (0.07-0.42) |
| Rollan, et al., 2002 <sup>54</sup>        | No known CAD | Fair    | 99 (women)   | ≥ 50   | 42           | 69 (55-83)             | 89 (82-97)             | 83 (70-95)     | 80 (70-90)     | 6.56 (3.00-14.36)  | 0.35 (0.22-0.55) |
| Williams, et al., 1994 <sup>74</sup>      | No known CAD | Fair    | 70 (women)   | ≥ 50   | 47           | 88 (77-99)             | 84 (72-96)             | 83 (70-95)     | 89 (78-99)     | 5.42 (2.58-11.40)  | 0.14 (0.06-0.37) |
| Marwick, et al., 1995 <sup>55</sup>       | No known CAD | Fair    | 161 (women)  | ≥ 50   | 37           | 80 (69-90)             | 81 (74-89)             | 71 (60-82)     | 87 (81-94)     | 4.28 (2.79-6.55)   | 0.25 (0.15-0.42) |
| Dionisopoulos, et al., 1997 <sup>75</sup> | No known CAD | Fair    | 101 (women)  | ≥ 50   | 66           | 90 (82-97)             | 79 (66-93)             | 90 (82-97)     | 79 (66-93)     | 4.35 (2.24-8.46)   | 0.13 (0.06-0.27) |
|   |              |         | 137 (men)    |        | 76           | 85 (78-91)             | 96 (85-99)             | 98 (96-100)    | 67 (56-79)     | 19.2 (4.94-74.44)  | 0.15 (0.10-0.23) |
| Mazeika, et al., 1992 <sup>76</sup>       | No known CAD | Fair    | 14 (women)   | ≥ 70   | 50           | 57 (20-94)             | 71 (38-100)            | 67 (29-100)    | 63 (29-96)     | 2.00 (0.53-7.60)   | 0.6 (0.23-1.60)  |
|   |              |         | 41 (men)     |        | 80           | 33 (18-52)             | 88 (47-1.00)           | 92 (76-100)    | 24 (9-40)      | 2.67 (0.40-17.76)  | 0.76 (0.53-1.09) |
| Slavich, et al., 1996 <sup>77</sup>       | No known CAD | Fair    | 49 (women)   | ≥ 50   | 45           | 59 (39-80)             | 75 (58-92)             | 68 (48-89)     | 67 (49-84)     | 2.36 (1.09-5.13)   | 0.55 (0.31-0.95) |
| Takeuchi, et al., 1996 <sup>78</sup>      | No known CAD | Fair    | 70 (women)   | ≥ 50   | 29           | 75 (56-94)             | 92 (84-100)            | 79 (61-97)     | 90 (82-98)     | 9.38 (3.54-24.82)  | 0.27 (0.13-0.58) |

**Table 3 Summary of accuracy data evaluating ECHO for diagnosing CAD (continued)**

| Study                                 | Patient Mix  | Quality | Patients (N) | Cath % | Prevalence % | Sensitivity % (95% CI) | Specificity % (95% CI) | PPV % (95% CI) | NPV % (95% CI) | LR+ (95% CI)       | LR- (95% CI)     |
|---------------------------------------|--------------|---------|--------------|--------|--------------|------------------------|------------------------|----------------|----------------|--------------------|------------------|
| Masini, et al., 1988 <sup>61</sup>    | No known CAD | Fair    | 68 (women)   | ≥ 70   | 38           | 77 (61-93)             | 93 (85-100)            | 87 (73-100)    | 87 (77-97)     | 10.77 (3.55-32.70) | 0.25 (0.12-0.50) |
| Roger, et al., 1997 <sup>79</sup>     | No known CAD | Poor    | 96 (women)   | ≥ 50   | 60           | 79 (69-90)             | 37 (22-52)             | 66 (55-77)     | 54 (35-73)     | 1.26 (0.95-1.66)   | 0.56 (0.29-1.08) |
|                                       |              |         | 244 (men)    |        | 80           | 78 (71-83)             | 44 (30-59)             | 84 (79-90)     | 34 (22-45)     | 1.39 (1.08-1.80)   | 0.50 (0.33-0.76) |
| Laurienzo, et al., 1997 <sup>39</sup> | Mixed        | Good    | 84 (women)   | ≥ 70   | 26           | 82 (66-98)             | 100 (95-100)           | 100 (83-100)   | 94 (88-100)    | NA                 | 0.18 (0.07-0.44) |
| Elhendy, et al., 1998 <sup>71</sup>   | Mixed        | Good    | 70 (women)   | ≥ 50   | 64           | 78 (66-90)             | 92 (81-100)            | 95 (87-100)    | 70 (54-85)     | 9.72 (2.55-37.07)  | 0.24 (0.14-0.42) |
| Lewis, et al., 1999 <sup>72</sup>     | Mixed        | Good    | 92 (women)   | ≥ 50   | 27           | 40 (21-59)             | 81 (71-90)             | 43 (23-64)     | 78 (69-88)     | 2.06 (1.04-4.09)   | 0.74 (0.53-1.05) |
| Shin, et al., 2003 <sup>69</sup>      | Mixed        | Fair    | 162 (women)  | ≥ 50   | 34           | 82 (72-92)             | 71 (62-80)             | 59 (48-70)     | 88 (82-95)     | 2.82 (2.05-3.90)   | 0.26 (0.14-0.45) |
|                                       |              |         | 302 (men)    |        | 55           | 79 (72-85)             | 80 (72-86)             | 83 (77-89)     | 76 (68-83)     | 3.95 (2.80-5.59)   | 0.26 (0.19-0.36) |
| Elhendy, et al., 1997 <sup>35</sup>   | Mixed        | Fair    | 96 (women)   | ≥ 50   | 65           | 76 (65-86)             | 94 (86-100)            | 96 (90-100)    | 68 (55-81)     | 12.89 (3.33-49.80) | 0.26 (0.16-0.40) |
|                                       |              |         | 210 (men)    |        | 81           | 73 (66-80)             | 77 (61-89)             | 93 (89-98)     | 39 (28-50)     | 3.17 (1.77-5.66)   | 0.35 (0.26-0.47) |
| Bjornstad, et al., 1995 <sup>70</sup> | Mixed        | Fair    | 7 (women)    | ≥ 50   | 86           | 83 (54-100)            | 100 (NA-100)           | 100 (40-100)   | 50 (NA-100)    | NA                 | 0.17 (0.03-1.00) |
|                                       |              |         | 30 (men)     |        | 93           | 68 (48-84)             | 100 (55-1.00)          | 100 (84-100)   | 36 (11-61)     | NA                 | 0.32 (0.19-0.55) |
| Agati, et al., 1992 <sup>56</sup>     | Mixed        | Fair    | 10 (women)   | ≥ 70   | 50           | 80 (45-100)            | 100 (40-100)           | 100 (25-100)   | 83 (54-100)    | NA                 | 0.20 (0.03-1.15) |
|                                       |              |         | 22 (men)     |        | 86           | 95 (74-1.00)           | 100 (37-1.00)          | 100 (83-100)   | 75 (33-100)    | NA                 | 0.05 (0.01-0.35) |
| Masini, et al., 1988 <sup>61</sup>    | Mixed        | Fair    | 83 (women)   | ≥ 70   | 47           | 79 (67-92)             | 93 (86-100)            | 91 (82-100)    | 84 (73-94)     | 11.66 (3.87-35.16) | 0.22 (0.12-0.41) |
| Ho, et al., 1998 <sup>67</sup>        | Mixed        | Fair    | 51 (women)   | ≥ 50   | 57           | 93 (77-99)             | 82 (60-95)             | 87 (75-99)     | 90 (77-100)    | 5.12 (2.10-12.49)  | 0.08 (0.02-0.32) |

Abbreviations: CAD = coronary artery disease; Cath % = % stenosis defined to be positive for CAD on diagnostic cardiac catheterization (coronary angiography); CI = confidence interval; LR+ = positive likelihood ratio; LR- = negative likelihood ratio; NPV = negative predictive value; NR = not reported; PPV = positive predictive value



## SPECT

We identified 30 studies evaluating the accuracy of SPECT compared with coronary angiography (Table 4).<sup>26,28,37,39,40,43,50,52,54,62,67,68,71,77,78,80-94</sup> Fourteen of these studies were conducted exclusively in women with no known CAD, and these are the studies used in our primary analysis. In our secondary analysis, we evaluated the accuracy of SPECT in diagnosing CAD in mixed populations of known and no known CAD, including 16 additional studies.

### Primary Analysis: Population of Women With No Known CAD

The 14 studies represent findings on SPECT use in 1000 women (sample size ranging from 19 to 184 women). Of these studies, four were good quality, nine were fair quality, and 1 was poor quality. Sensitivity varied from 62 to 93 percent, and specificity varied from 50 to 91 percent; the median sensitivity was 82 percent, and the median specificity was 81 percent. Figure 20 presents forest plots of the individual study estimates of sensitivity and specificity of SPECT for diagnosing CAD in women with no known CAD.

**Figure 20. Accuracy of SPECT in women with no known CAD**

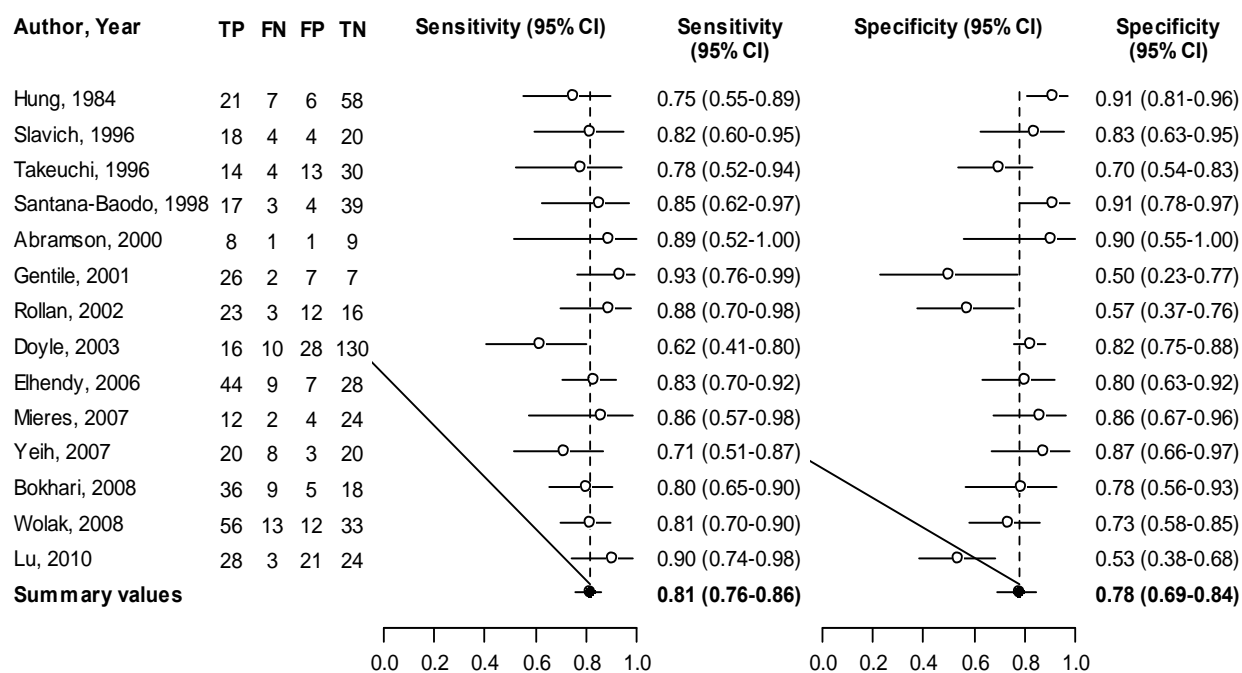
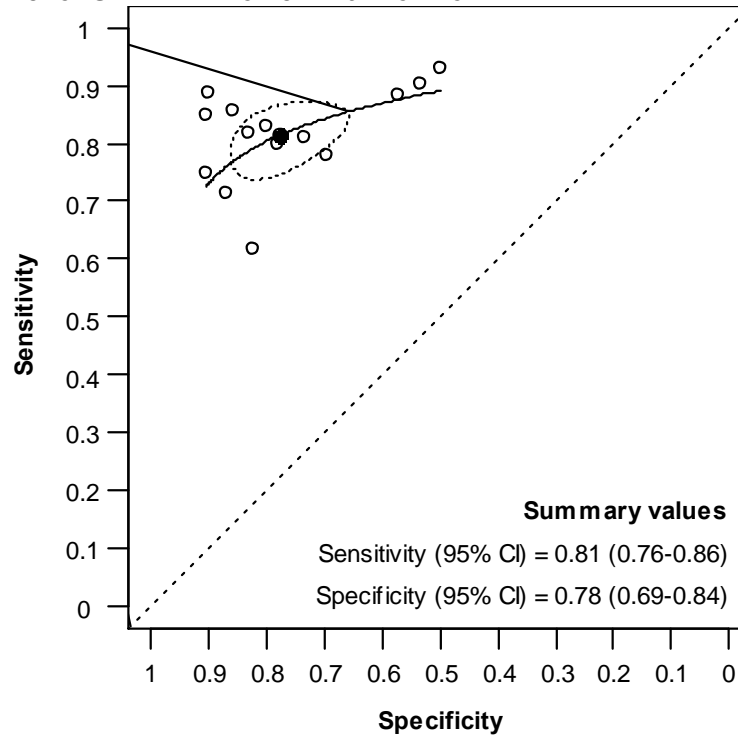


Figure 21 presents an SROC curve with an average sensitivity of 81 percent (95% CI, 76 to 86 percent) and specificity of 78 percent (95% CI, 69 to 84 percent).

**Figure 21. SROC curve for SPECT in women with no known CAD**



The prevalence of CAD in these 14 studies ranged from 14 to 67 percent with a mean prevalence of 45 percent. In the individual studies, PPV ranged from 36 to 89 percent, and NPV ranged from 50 to 91 percent. LR+ ranged from 1.86 to 9.14 and LR- from 0.12 to 0.47 respectively. Using the summary sensitivity and specificity of 81 and 78 percent, respectively, we calculated an overall PPV of 75 percent and a negative predictive value of 83 percent. Similarly, we calculated summary LR+ of 3.68 and LR- of 0.24.

### **Accuracy of SPECT in Four Good-Quality Studies**

Next, we evaluated the accuracy of SPECT compared with coronary angiography in the four good-quality studies. In these studies, sensitivity varied from 62 to 93 percent, and specificity varied from 50 to 91 percent; the median sensitivity was 83 percent, and the median specificity was 68 percent. Figure 22 presents forest plots of the individual study estimates of sensitivity and specificity of SPECT for diagnosing CAD in women with no known CAD.

**Figure 22. Accuracy of SPECT in four good-quality studies in women with no known CAD**

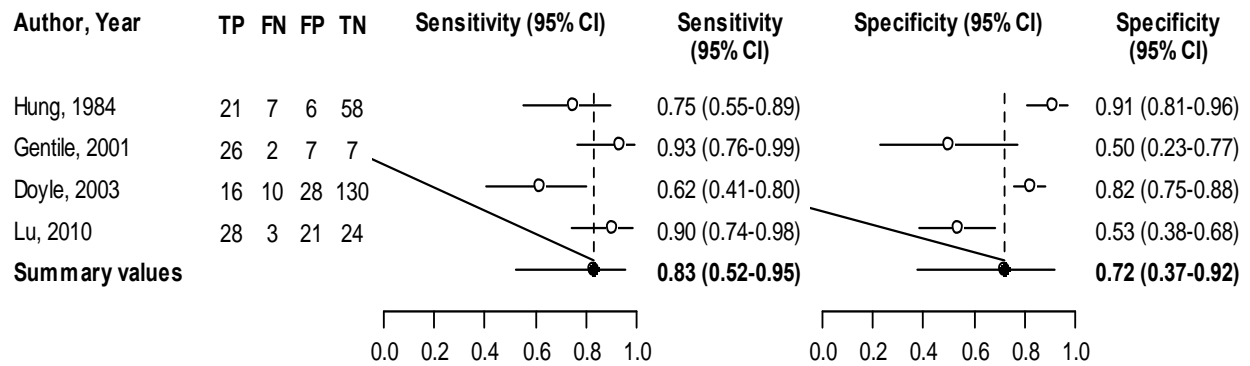
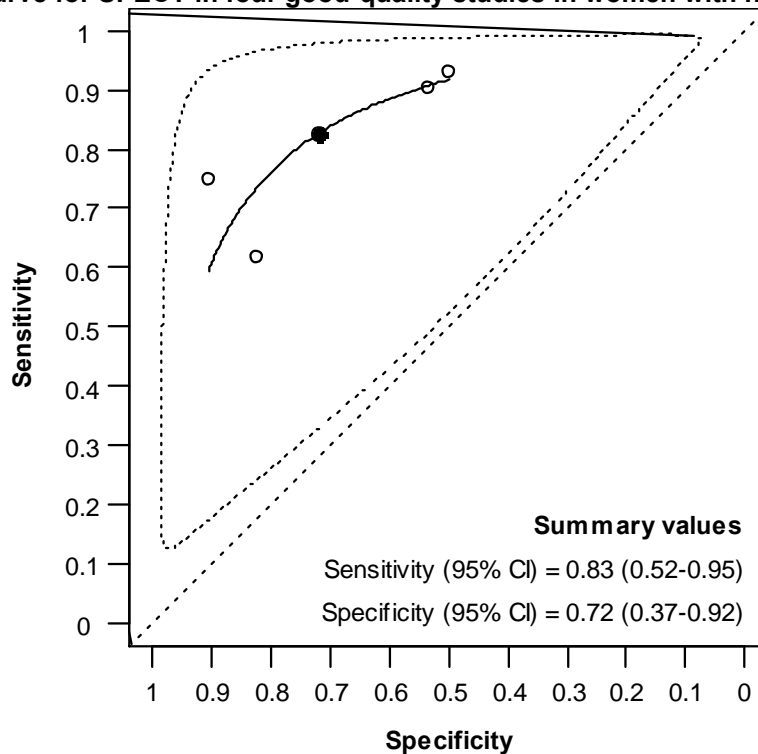


Figure 23 presents an SROC curve with an average sensitivity of 83 percent (95% CI, 52 to 95 percent) and specificity of 72 percent (95% CI, 37 to 92 percent). It is important to note that given the small number of studies and wide confidence intervals that these summary statistics should be interpreted with caution.

**Figure 23. SROC curve for SPECT in four good-quality studies in women with no known CAD**



The prevalence of CAD in these 4 good-quality studies ranged from 14 to 67 percent with a mean prevalence of 38 percent. In the individual studies, PPV ranged from 36 to 79 percent, and NPV ranged from 78 to 93 percent. LR+ ranged from 1.86 to 8.0 and LR- from 0.14 to 0.47. Using the summary sensitivity and specificity of 83 and 72 percent, respectively, we calculated an overall PPV of 65 percent and NPV of 88 percent. Similarly, we calculated summary LR+ of 2.96 and LR- ratio of 0.24.

### **Secondary Analysis: Mixed Population of Women With Known and No Known CAD**

We performed a secondary analysis where we expanded our inclusion criteria to include studies whose patient population included a mix of women with known CAD and women with no known CAD. This expanded inclusion criteria allowed an additional 16 studies to be included in our analysis (totaling 30 studies). The 30 studies represent findings on SPECT use in 2157 women (sample size ranging from 14 to 243 women). Of these 30 studies, 10 were good quality, 15 were fair quality, and 5 were poor quality (Table 4).

In these 30 studies, sensitivity varied from 15 to 100 percent, and specificity varied from 27 to 100 percent; the median sensitivity was 83 percent, and the median specificity was 81 percent. Figure 24 presents forest plots of the individual study estimates of sensitivity and specificity of SPECT for diagnosing CAD in women from mixed populations.

**Figure 24. Accuracy of SPECT in women from mixed populations**

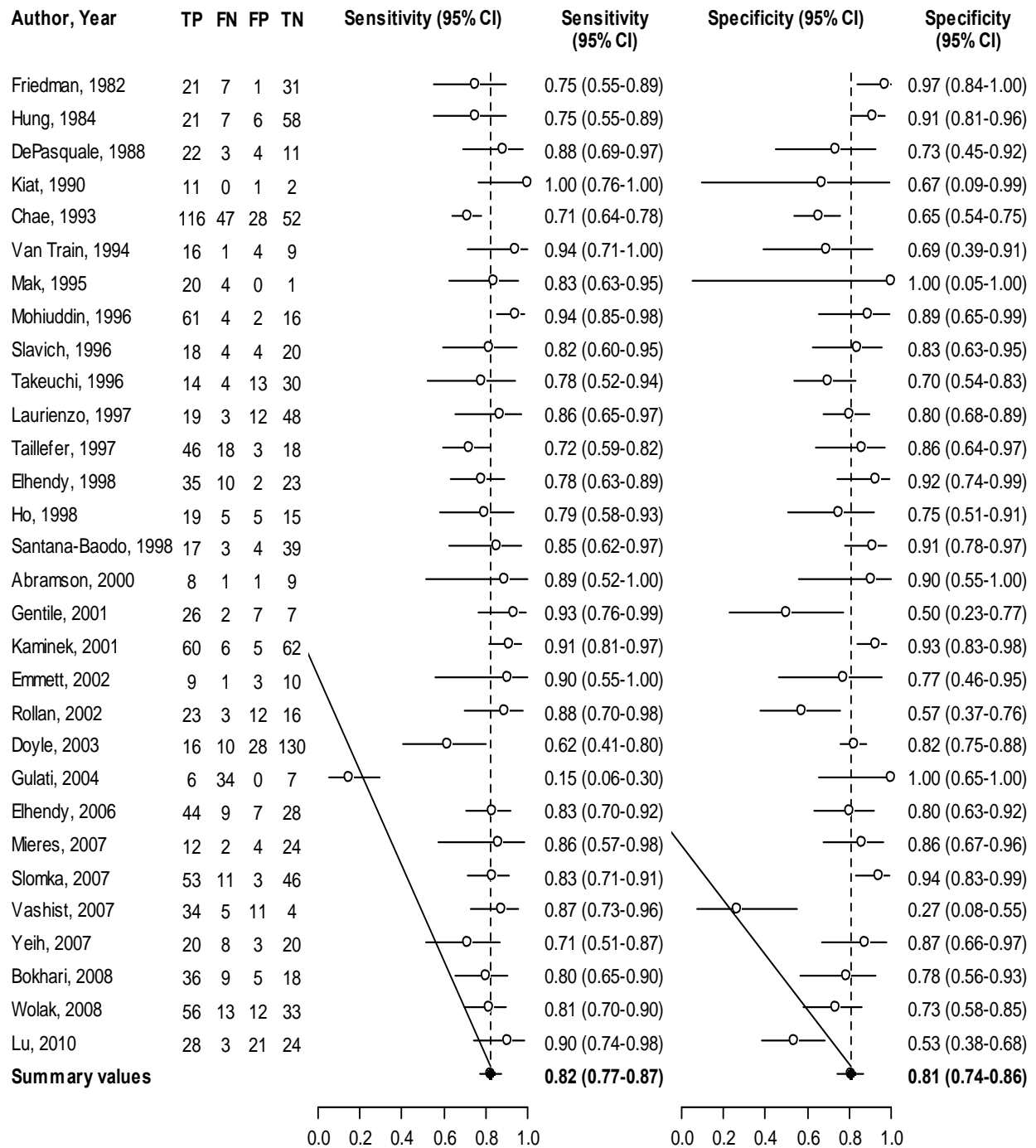
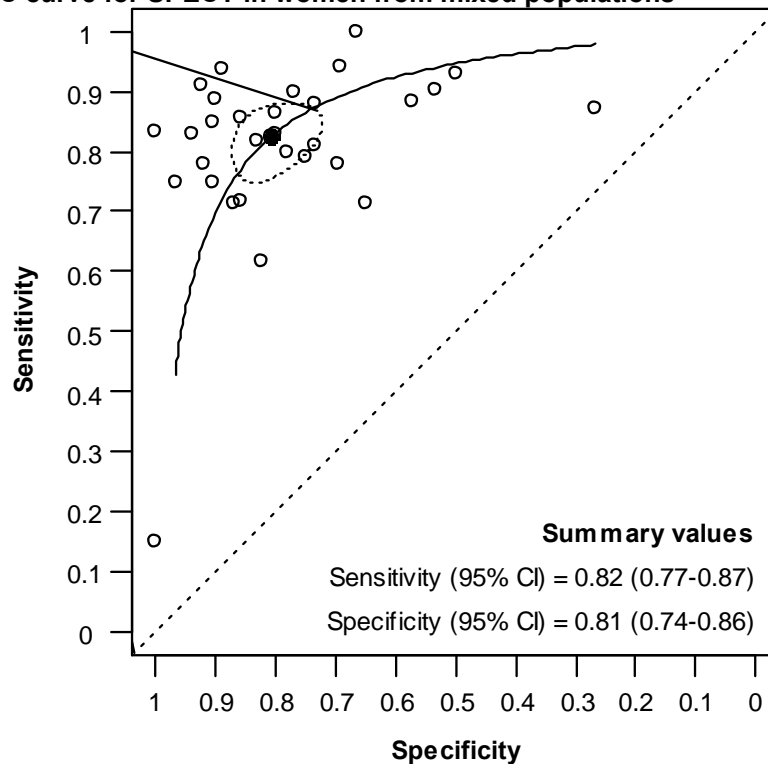


Figure 25 presents an SROC curve demonstrating an average sensitivity of 82 percent (95% CI, 77 to 87 percent) and specificity of 81 percent (95% CI, 74 to 86 percent).

**Figure 25. SROC curve for SPECT in women from mixed populations**



The prevalence of CAD in the 30 studies ranged from 14 to 96 percent with a mean prevalence of 54 percent. In the individual studies, PPV ranged from 36 to 100 percent, and NPV ranged from 27 to 100 percent. LR+ ranged from 1.89 to 24 and LR- from 0 to 0.89. Using the summary sensitivity and specificity of 82 and 81 percent, respectively, we calculated an overall PPV of 84 percent and NPV of 79 percent. Similarly, we calculated summary LR+ of 4.32 and LR- of 0.22.

### **Accuracy of SPECT in 10 Good-Quality Studies**

Next, we evaluated the accuracy of SPECT compared with coronary angiography in the 10 good-quality studies. In these studies, sensitivity varied from 62 to 94 percent, and specificity varied from 50 to 100 percent; the median sensitivity was 81 percent, and the median specificity was 84 percent. Figure 26 presents forest plots of the individual study estimates of sensitivity and specificity of SPECT for diagnosing CAD in women with from mixed populations.

**Figure 26. Accuracy of SPECT in 10 good-quality studies in women from mixed populations**

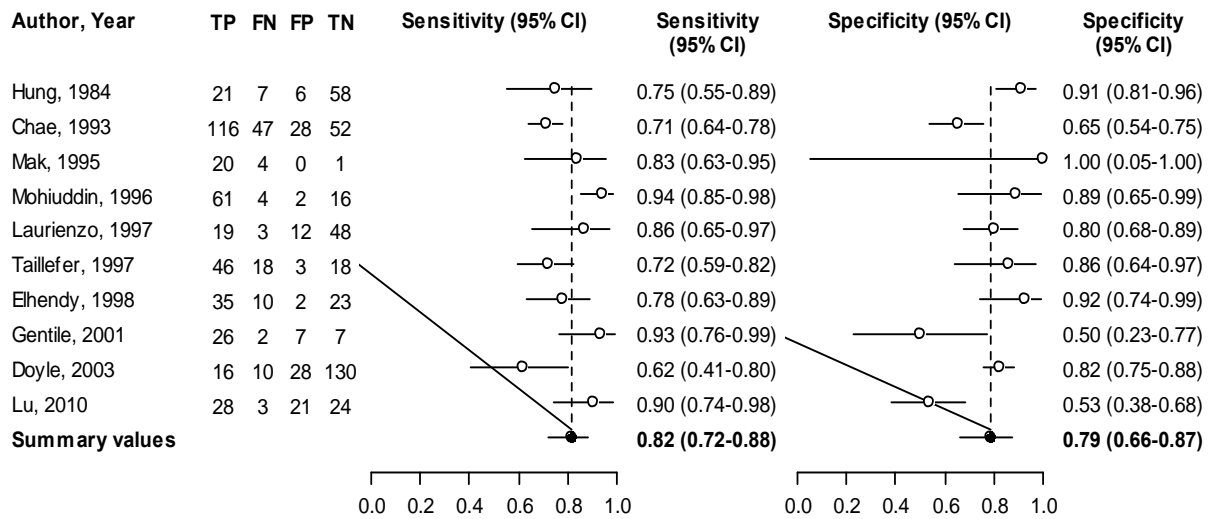
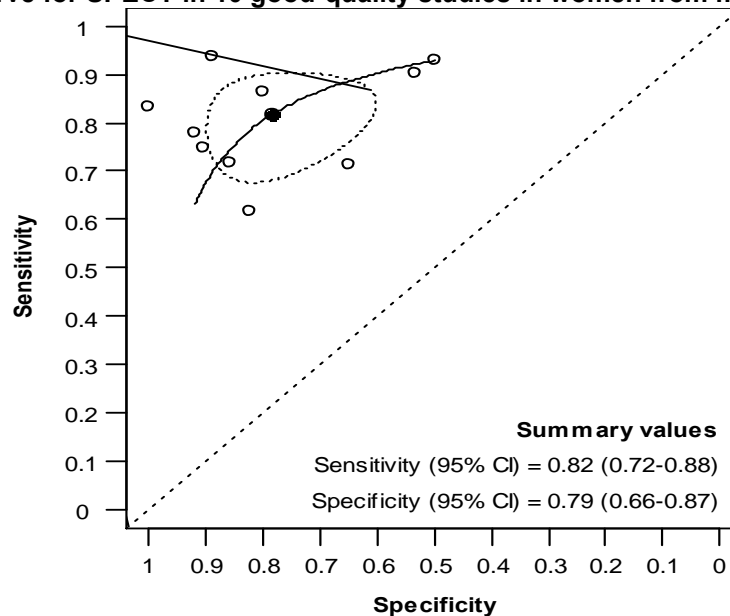


Figure 27 presents an SROC curve demonstrating an average sensitivity of 82 percent (95% CI, 72 to 88 percent) and specificity of 79 percent (95% CI, 66 to 87 percent).

**Figure 27. SROC curve for SPECT in 10 good-quality studies in women from mixed populations**



The prevalence of CAD in these 10 good-quality studies ranged from 14 to 96 percent with a mean prevalence of 56 percent. In the individual studies, PPV ranged from 36 to 100 percent, and NPV ranged from 20 to 94 percent. LR+ ranged from 1.86 to 9.72 and LR- from 0.07 to 0.47. Using the summary sensitivity and specificity of 82 and 79 percent, respectively, we calculated an overall PPV of 84 percent and NPV of 78 percent. Similarly, we calculated summary LR+ of 3.90 and LR- of 0.23.

**Table 4. Summary of accuracy data evaluating SPECT for diagnosing CAD**

| Study                                     | Patient Mix  | Quality | Patients (N) | Cath % | Prevalence % | Sensitivity % (95% CI) | Specificity % (95% CI) | PPV % (95% CI) | NPV % (95% CI) | LR+ (95% CI)      | LR- (95% CI)     |
|---|--------------|---------|--------------|--------|--------------|------------------------|------------------------|----------------|----------------|-------------------|------------------|
| Lu, et al., 2010 <sup>28</sup>            | No known CAD | Good    | 76 (women)   | ≥ 50   | 41           | 90 (74-98)             | 53 (38-68)             | 57 (43-71)     | 89 (77-100)    | 1.94 (1.39-2.70)  | 0.18 (0.06-0.55) |
| Doyle, et al., 2003 <sup>26</sup>         | No known CAD | Good    | 184 (women)  | ≥ 70   | 14           | 62 (41-80)             | 82 (75-88)             | 36 (22-51)     | 93 (89-97)     | 3.47 (2.21-5.46)  | 0.47 (0.29-0.76) |
| Gentile, et al., 2001 <sup>43</sup>       | No known CAD | Good    | 42 (women)   | ≥ 60   | 67           | 93 (76-99)             | 50 (23-77)             | 79 (65-93)     | 78 (51-100)    | 1.86 (1.09-3.17)  | 0.14 (0.03-0.60) |
|   |              |         | 90 (men)     |        | 89           | 94 (86-98)             | 60 (26-88)             | 95 (90-100)    | 55 (25-84)     | 2.34 (1.09-5.02)  | 0.10 (0.04-0.28) |
| Hung, et al., 1984 <sup>90</sup>          | No known CAD | Good    | 92 (women)   | ≥ 70   | 30           | 75 (55-89)             | 91 (81-96)             | 78 (62-93)     | 89 (82-97)     | 8 (3.63-17.6)     | 0.28 (0.14-0.53) |
| Wolak, et al., 2008 <sup>91</sup>         | No known CAD | Fair    | 114 (women)  | ≥ 70   | 61           | 81 (70-90)             | 73 (58-85)             | 82 (73-91)     | 72 (59-85)     | 3.04 (1.85-5.01)  | 0.26 (0.15-0.43) |
| Yeih, et al., 2007 <sup>50</sup>          | No known CAD | Fair    | 51 (women)   | ≥ 50   | 55           | 71 (51-87)             | 87 (66-97)             | 87 (73-100)    | 71 (55-88)     | 5.48 (1.86-16.14) | 0.33 (0.18-0.60) |
| Mieres, et al., 2007 <sup>52</sup>        | No known CAD | Fair    | 42 (women)   | ≥ 50   | 33           | 86 (57-98)             | 86 (67-96)             | 75 (54-96)     | 92 (82-100)    | 6 (2.36-15.24)    | 0.17 (0.05-0.61) |
| Elhendy, et al., 2006 <sup>92</sup>       | No known CAD | Fair    | 88 (women)   | ≥ 50   | 60           | 83 (70-92)             | 80 (63-92)             | 86 (77-96)     | 76 (62-90)     | 4.15 (2.12-8.14)  | 0.21 (0.11-0.39) |
| Rollan, et al., 2002 <sup>54</sup>        | No known CAD | Fair    | 54 (women)   | ≥ 50   | 48           | 88 (70-98)             | 57 (37-76)             | 66 (50-81)     | 84 (68-100)    | 2.06 (1.32-3.24)  | 0.20 (0.07-0.61) |
| Abramson, et al., 2000 <sup>93</sup>      | No known CAD | Fair    | 19 (women)   | ≥ 50   | 47           | 89 (52-100)            | 90 (55-100)            | 89 (68-100)    | 90 (71-100)    | 8.89 (1.36-57.89) | 0.12 (0.02-0.8)  |
| Slavich, et al., 1996 <sup>77</sup>       | No known CAD | Fair    | 46 (women)   | ≥ 50   | 48           | 82 (60-95)             | 83 (63-95)             | 82 (66-98)     | 83 (68-98)     | 4.91 (1.96-12.27) | 0.22 (0.09-0.54) |
| Takeuchi, et al., 1996 <sup>78</sup>      | No known CAD | Fair    | 61 (women)   | ≥ 50   | 30           | 78 (52-94)             | 70 (54-83)             | 52 (33-71)     | 88 (77-99)     | 2.57 (1.53-4.31)  | 0.32 (0.13-0.77) |
| Santana-Baodo, et al., 1998 <sup>68</sup> | No known CAD | Fair    | 63 (women)   | ≥ 50   | 32           | 85 (62-97)             | 91 (78-97)             | 81 (64-98)     | 93 (85-100)    | 9.13 (3.53-23.65) | 0.16 (0.06-0.47) |
|   |              |         | 100 (men)    |        | 80           | 92 (84-97)             | 90 (68-99)             | 97 (94-100)    | 75 (58-92)     | 9.25 (2.48-34.5)  | 0.08 (0.04-0.18) |



**Table 4. Summary of accuracy data evaluating SPECT for diagnosing CAD (continued)**

| Study                                 | Patient Mix  | Quality | Patients (N) | Cath % | Prevalence % | Sensitivity %<br>(95% CI) | Specificity %<br>(95% CI) | PPV %<br>(95% CI) | NPV %<br>(95% CI) | LR+<br>(95% CI)      | LR-<br>(95% CI)     |
|---------------------------------------|--------------|---------|--------------|--------|--------------|---------------------------|---------------------------|-------------------|-------------------|----------------------|---------------------|
| Bokhari, et al., 2008 <sup>62</sup>   | No known CAD | Poor    | 68 (women)   | ≥ 50   | 66           | 80<br>(65-90)             | 78<br>(56-93)             | 88<br>(78-98)     | 67<br>(49-84)     | 3.68<br>(1.67-8.10)  | 0.26<br>(0.14-0.48) |
|                                       |              |         | 150 (men)    |        | 65           | 82<br>(73-89)             | 79<br>(65-89)             | 88<br>(81-95)     | 69<br>(58-81)     | 3.86<br>(2.26-6.58)  | 0.23<br>(0.15-0.36) |
| Laurienzo, et al., 1997 <sup>39</sup> | Mixed        | Good    | 84 (women)   | > 70   | 26           | 86<br>(65-97)             | 80<br>(68-89)             | 61<br>(44-78)     | 94<br>(88-100)    | 4.32<br>(2.54-7.36)  | 0.17<br>(0.06-0.49) |
| Taillefer, et al., 1997 <sup>87</sup> | Mixed        | Good    | 85 (women)   | ≥ 50   | 75           | 72<br>(59-82)             | 86<br>(64-97)             | 94<br>(87-100)    | 50<br>(34-66)     | 5.03<br>(1.75-14.50) | 0.33<br>(0.21-0.50) |
| Mak, et al., 1995 <sup>88</sup>       | Mixed        | Good    | 25 (women)   | ≥ 50   | 96           | 83<br>(63-95)             | 100<br>(5-100)            | 100<br>(85-100)   | 20<br>(15-55)     | NA                   | 0.17<br>(0.07-0.41) |
|                                       |              |         | 114 (men)    |        | 86           | 93<br>(86-97)             | 63<br>(35-85)             | 94<br>(89-99)     | 59<br>(35-82)     | 2.48<br>(1.31-4.67)  | 0.11<br>(0.05-0.26) |
| Chae, et al., 1993 <sup>40</sup>      | Mixed        | Good    | 243 (women)  | ≥ 50   | 67           | 71<br>(64-78)             | 65<br>(54-75)             | 81<br>(74-87)     | 53<br>(43-62)     | 2.03<br>(1.49-2.78)  | 0.44<br>(0.33-0.59) |
| Elhendy, et al., 1998 <sup>71</sup>   | Mixed        | Good    | 70 (women)   | ≥ 50   | 64           | 78<br>(63-89)             | 92<br>(74-99)             | 95<br>(87-100)    | 70<br>(54-85)     | 9.72<br>(2.55-37.07) | 0.24<br>(0.14-0.42) |
| Mohiuddin, et al., 1996 <sup>89</sup> | Mixed        | Good    | 83 (women)   | ≥ 50   | 78           | 94<br>(85-98)             | 89<br>(65-99)             | 97<br>(92-100)    | 80<br>(62-98)     | 8.45<br>(2.28-31.25) | 0.07<br>(0.03-0.18) |
|                                       |              |         | 119 (men)    |        | 80           | 87<br>(79-93)             | 83<br>(63-95)             | 95<br>(91-100)    | 63<br>(46-79)     | 5.24<br>(2.14-12.87) | 0.15<br>(0.09-0.26) |
| Vashist, et al., 2007 <sup>83</sup>   | Mixed        | Fair    | 54 (women)   | ≥ 50   | 72           | 87<br>(73-96)             | 27<br>(8-55)              | 76<br>(63-88)     | 44<br>(12-77)     | 1.19<br>(0.86-1.65)  | 0.48<br>(0.15-1.55) |
| Gulati, et al., 2004 <sup>84</sup>    | Mixed        | Fair    | 47 (women)   | ≥ 50   | 85           | 15<br>(6-30)              | 95<br>(65-100)            | 95<br>(78-100)    | 16<br>(5-28)      | 3.28<br>(.10-103.69) | 0.89<br>(0.72-1.10) |
|                                       |              |         | 25 (men)     |        | 96           | 4<br>(0-21)               | 100<br>(5-100)            | 99<br>(87-100)    | 4<br>(4-12)       | 7.39<br>(NA)         | 0.95<br>(0.77-1.16) |
| Emmett, et al., 2002 <sup>85</sup>    | Mixed        | Fair    | 23 (women)   | > 70   | 43           | 90<br>(55-100)            | 77<br>(46-95)             | 75<br>(51-100)    | 91<br>(74-100)    | 3.90<br>(1.42-10.75) | 0.13<br>(0.02-0.85) |
|                                       |              |         | 77 (men)     |        | 78           | 88<br>(77-95)             | 53<br>(28-77)             | 87<br>(78-95)     | 56<br>(32-81)     | 1.88<br>(1.12-3.13)  | 0.22<br>(0.10-0.50) |

**Table 4. Summary of accuracy data evaluating SPECT for diagnosing CAD (continued)**

| Study                                  | Patient Mix | Quality | Patients (N) | Cath % | Prevalence % | Sensitivity %<br>(95% CI) | Specificity %<br>(95% CI) | PPV %<br>(95% CI) | NPV %<br>(95% CI) | LR+<br>(95% CI)       | LR-<br>(95% CI)      |
|--|-------------|---------|--------------|--------|--------------|---------------------------|---------------------------|-------------------|-------------------|-----------------------|----------------------|
| Kiat, et al., 1990 <sup>86</sup>       | Mixed       | Fair    | 14 (women)   | ≥ 50   | 79           | 100<br>(76-100)           | 67<br>(9-99)              | 92<br>(76-100)    | 100<br>(50-100)   | 3<br>(0.61-14.86)     | 0<br>(NA)            |
|  |             |         | 39 (men)     |        | 95           | 92<br>(78-98)             | 50<br>(1-99)              | 97<br>(92-100)    | 25<br>(17-67)     | 1.84<br>(0.46-7.37)   | 0.16<br>(0.03-0.94)  |
| Friedman, et al., 1982 <sup>37</sup>   | Mixed       | Fair    | 60 (women)   | ≥ 70   | 47           | 75<br>(55-89)             | 97<br>(84-100)            | 95<br>(87-100)    | 82<br>(69-94)     | 24<br>(3.45-167.17)   | 0.26<br>(0.14-0.49)  |
| Ho, et al., 1998 <sup>67</sup>         | Mixed       | Fair    | 44 (women)   | ≥ 50   | 55           | 79<br>(58-93)             | 75<br>(51-91)             | 79<br>(63-95)     | 75<br>(56-94)     | 3.17<br>(1.44-6.95)   | 0.28<br>(0.12-0.63)  |
| Slomka, et al., 2007 <sup>80</sup>     | Mixed       | Poor    | 113 (women)  | ≥ 70   | 57           | 83<br>(71-91)             | 94<br>(83-99)             | 95<br>(89-100)    | 81<br>(70-91)     | 13.53<br>(4.49-40.71) | 0.18<br>(0.11-0.31)  |
| Kaminek, et al., 2001 <sup>81</sup>    | Mixed       | Poor    | 133 (women)  | ≥ 50   | 50           | 91<br>(81-97)             | 93<br>(83-98)             | 92<br>(86-99)     | 91<br>(84-98)     | 12.18<br>(5.22-28.41) | 0.10<br>(0.046-0.21) |
|  |             |         | 455 (men)    |        | 83           | 94<br>(91-96)             | 82<br>(71-90)             | 96<br>(94-98)     | 72<br>(63-82)     | 5.15<br>(3.20-8.28)   | 0.08<br>(0.05-0.12)  |
| DePasquale, et al., 1988 <sup>82</sup> | Mixed       | Poor    | 40 (women)   | ≥ 70   | 63           | 88<br>(69-97)             | 73<br>(45-92)             | 85<br>(71-98)     | 79<br>(57-100)    | 3.30<br>(1.41-7.73)   | 0.16<br>(0.05-0.49)  |
|  |             |         | 170 (men)    |        | 91           | 96<br>(92-99)             | 25<br>(7-52)              | 93<br>(88-97)     | 40<br>(10-70)     | 1.28<br>(0.96-1.70)   | 0.16<br>(0.05-0.49)  |
| Van Train, et al., 1994 <sup>94</sup>  | Mixed       | Poor    | 39 (women)   | ≥ 50   | 44           | 94<br>(71-100)            | 69<br>(39-91)             | 80<br>(62-98)     | 90<br>(71-100)    | 3.06<br>(1.34-6.97)   | 0.08<br>(0.01-0.59)  |
|  |             |         | 94 (men)     |        | 90           | 89<br>(81-95)             | 44<br>(14-79)             | 94<br>(89-99)     | 31<br>(6-56)      | 1.61<br>(0.89-2.90)   | 0.24<br>(0.09-0.62)  |

Abbreviations: CAD = coronary artery disease; Cath % = % stenosis defined to be positive for CAD on diagnostic cardiac catheterization (coronary angiography); CI = confidence interval; LR+ = positive likelihood ratio; LR- = negative likelihood ratio; NPV = negative predictive value; NR = not reported; PPV = positive predictive value

## CMR

We identified six studies evaluating the accuracy of CMR compared with coronary angiography (Table 5).<sup>26,95-99</sup> Five of these studies reported accuracy data in women with no known CAD, and these are the studies used in our primary analysis. In our secondary analysis, we evaluated the accuracy of CMR in diagnosing CAD in mixed populations of known and no known CAD, including additional data from two studies<sup>95,96</sup> and data from one additional study.<sup>99</sup>

### Primary Analysis: Population of Women With No Known CAD

The five studies represent findings on CMR use in 501 women (sample size ranging from 30 to 184 women). All five of these studies were rated good quality. In these studies, sensitivity varied from 58 to 83 percent, and specificity varied from 59 to 96 percent; the median sensitivity was 75 percent, and the median specificity was 88 percent. Figure 28 presents forest plots of the individual study estimates of sensitivity and specificity of CMR for diagnosing CAD in women with no known CAD.

**Figure 28. Accuracy of CMR in women with no known CAD**

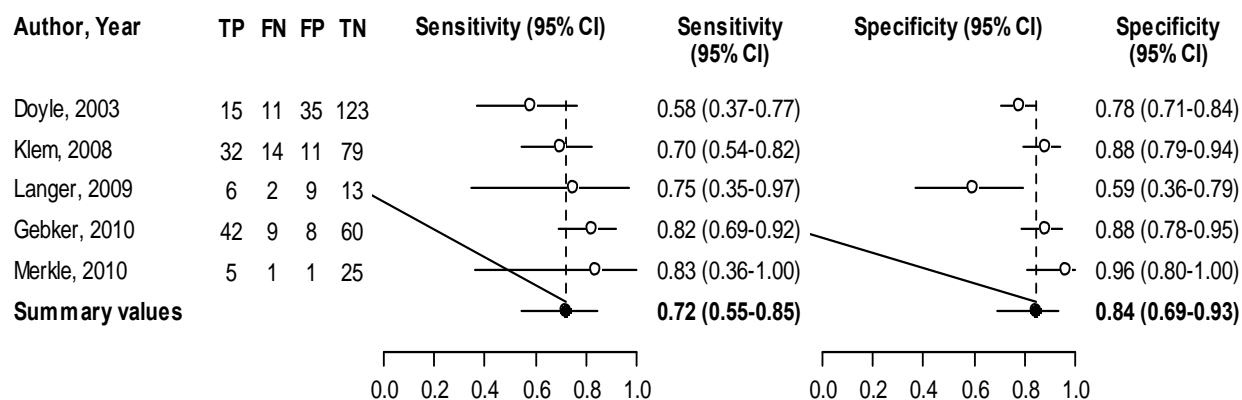
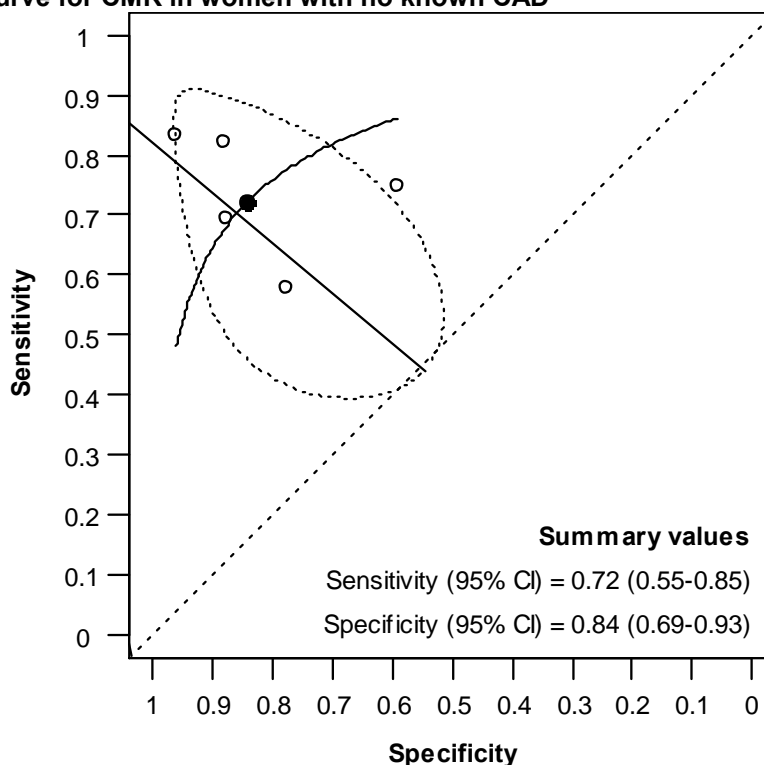


Figure 29 presents an SROC curve with an average sensitivity of 72 percent (95% CI, 55 to 85 percent) and specificity of 84 percent (95% CI, 69 to 93 percent).

**Figure 29. SROC curve for CMR in women with no known CAD**



The prevalence of CAD in these 5 studies ranged from 14 to 43 percent with a mean prevalence of 27 percent. In the individual studies, PPV ranged from 30 to 84 percent, and NPV ranged from 59 to 96 percent. LR+ ranged from 1.83 to 21.67 and LR- from 0.17 to 0.54. Using the summary sensitivity and specificity of 72 and 84 percent, respectively, we calculated an overall PPV of 62 percent and NPV of 89 percent. Similarly, we calculated summary LR+ of 4.5 and LR- of 0.33.

### **Secondary Analysis: Mixed Population of Women With Known and No Known CAD**

We performed a secondary analysis where we expanded our inclusion criteria to include studies whose patient population included a mix of women with known CAD and women with no known CAD. This expanded inclusion criteria allowed an additional 64 patients from one study<sup>95</sup> and an additional 45 patients from another study<sup>96</sup> to be included in our analysis—as well as 168 patients from a third study that was not included in our primary analysis<sup>99</sup> (totaling 6 studies). The 6 studies represent findings on CMR use in 778 women (sample size ranging from 30 to 184 women). Five of these studies were good-quality, and one was fair quality (Table 5).

In these 6 studies, sensitivity varied from 58 to 92 percent, and specificity varied from 59 to 91 percent; the median sensitivity was 80 percent, and the median specificity was 83 percent. Figure 30 presents forest plots of the individual study estimates of sensitivity and specificity of CMR for diagnosing CAD in women from mixed populations.

**Figure 30. Accuracy of CMR in women from mixed populations**

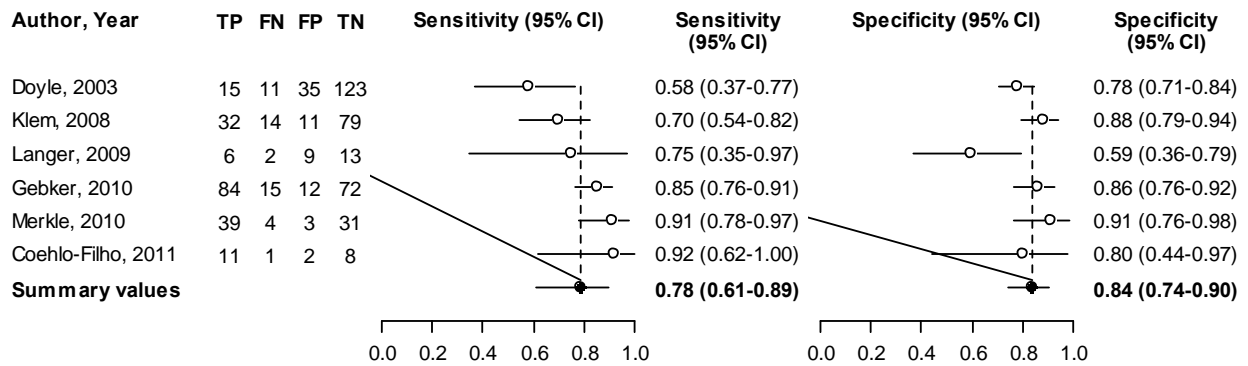
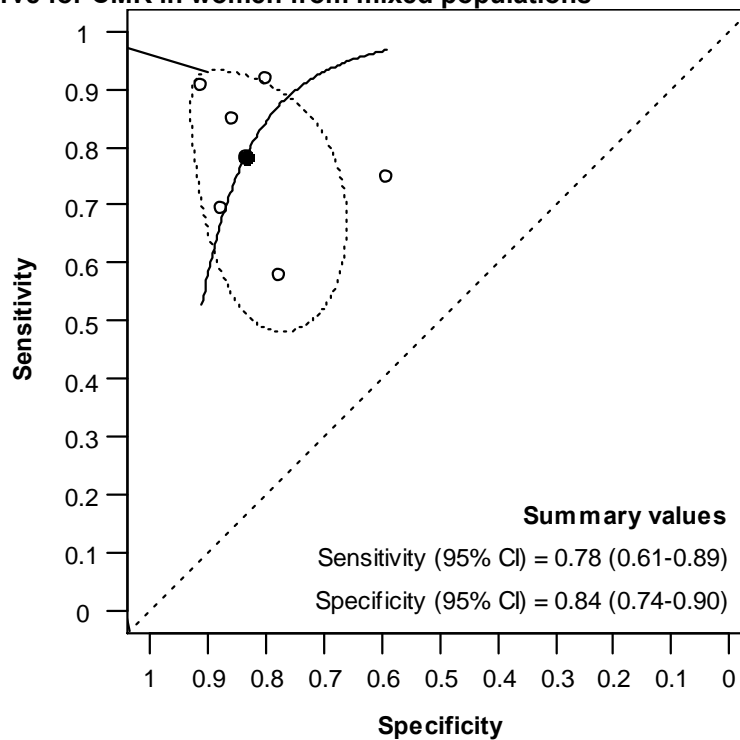


Figure 31 presents an SROC curve demonstrating an average sensitivity of 78 percent (95% CI, 61 to 89 percent) and specificity of 84 percent (95% CI, 74 to 90 percent).

**Figure 31. SROC curve for CMR in women from mixed populations**



The prevalence of CAD in the 6 studies ranged from 7 to 56 percent with a mean prevalence of 32 percent. In the individual studies, PPV ranged from 30 to 93 percent, and NPV ranged from 59 to 91 percent. LR+ ranged from 1.83 to 10.28 and LR- from 0.102 to 0.54. Using the summary sensitivity and specificity of 78 and 84 percent, respectively, we calculated an overall PPV of 69 percent and NPV of 89 percent. Similarly, we calculated summary LR+ of 4.88 and LR- of 0.26.

### Accuracy of CMR in Five Good-quality Studies

Next, we evaluated the accuracy of CMR compared with coronary angiography in the five good-quality studies. In these studies, sensitivity varied from 58 to 91 percent, and specificity varied from 59 to 91 percent; the median sensitivity was 75 percent, and the median specificity was 86 percent. Figure 32 presents forest plots of the individual study estimates of sensitivity and specificity of CMR for diagnosing CAD in women from mixed populations.

**Figure 32. Accuracy of CMR in five good-quality studies in women from mixed populations**

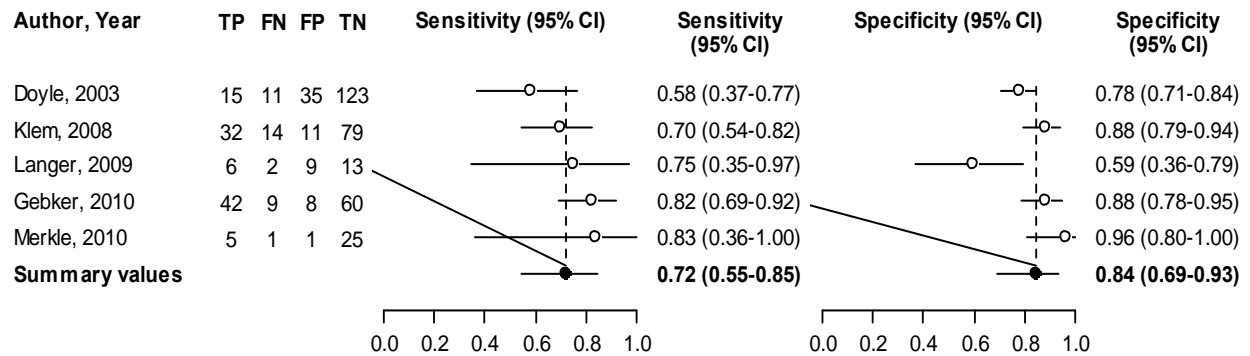
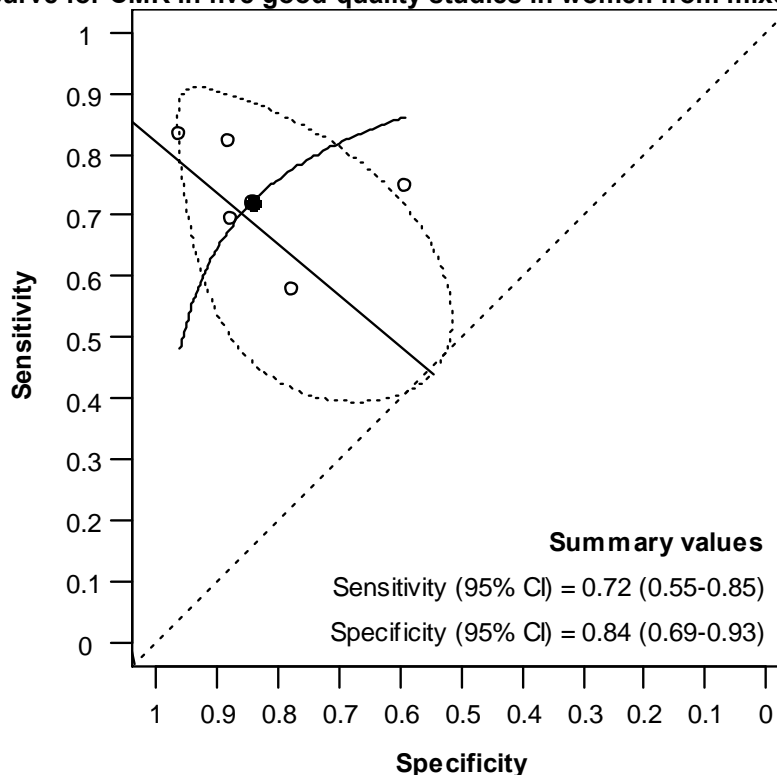


Figure 33 presents an SROC curve demonstrating an average sensitivity of 72 percent (95% CI, 55 to 85 percent) and specificity of 84 percent (95% CI, 69 to 93 percent).

**Figure 33. SROC curve for CMR in five good-quality studies in women from mixed populations**



The prevalence of CAD in these five good-quality studies ranged from 14 to 56 percent with a mean prevalence of 37 percent. In the individual studies, PPV ranged from 30 to 93 percent, and NPV ranged from 59 to 91 percent. LR+ ranged from 1.83 to 10.28 and LR- from 0.102 to 0.54. Using the summary sensitivity and specificity of 72 and 84 percent, respectively, we calculated an overall PPV of 72 percent and NPV of 84 percent. Similarly, we calculated summary LR+ of 4.5 and LR- of 0.33.

**Table 5. Summary of accuracy data evaluating CMR for diagnosing CAD**

| Study                                    | Patient Mix  | Quality | Patients (N) | Cath % | Prevalence % | Sensitivity % (95% CI) | Specificity % (95% CI) | PPV % (95% CI) | NPV % (95% CI) | LR+ (95% CI)        | LR- (95% CI)     |
|--|--------------|---------|--------------|--------|--------------|------------------------|------------------------|----------------|----------------|---------------------|------------------|
| Merkle, et al., 2010 <sup>96</sup>       | No known CAD | Good    | 32 (women)   | ≥ 50   | 19           | 83 (36-100)            | 96 (80-100)            | 83 (54-100)    | 96 (89-100)    | 21.67 (3.07-153.05) | 0.17 (0.03-1.04) |
|  |              |         | 41 (men)     |        | 34           | 79 (57-100)            | 74 (58-91)             | 61 (39-84)     | 87 (73-100)    | 3.03 (1.51-6.07)    | 0.29 (0.10-0.81) |
| Klem, et al., 2008 <sup>97</sup>         | No known CAD | Good    | 136 (women)  | ≥ 50   | 34           | 70 (54-82)             | 88 (79-94)             | 74 (61-87)     | 85 (78-92)     | 5.69 (3.17-10.22)   | 0.35 (0.22-0.54) |
| Langer, et al., 2009 <sup>98</sup>       | No known CAD | Good    | 30 (women)   | ≥ 50   | 27           | 75 (35-97)             | 59 (36-79)             | 40 (15-65)     | 87 (69-100)    | 1.83 (0.96-3.48)    | 0.42 (0.12-1.48) |
|  |              |         | 38 (men)     |        | 47           | 67 (45-88)             | 70 (50-90)             | 67 (45-88)     | 70 (50-90)     | 2.22 (1.06-4.68)    | 0.48 (0.23-0.97) |
| Doyle, et al., 2003 <sup>26</sup>        | No known CAD | Good    | 184 (women)  | ≥ 70   | 14           | 58 (37-77)             | 78 (71-84)             | 30 (17-43)     | 92 (87-96)     | 2.60 (1.68-4.04)    | 0.54 (0.34-0.86) |
| Gebker, et al., 2010 <sup>95</sup>       | No known CAD | Good    | 119 (women)  | ≥ 70   | 43           | 82 (69-92)             | 88 (78-95)             | 84 (74-94)     | 87 (79-95)     | 7.00 (3.61-13.59)   | 0.20 (0.11-0.36) |
| Gebker, et al., 2010 <sup>95</sup>       | Mixed        | Good    | 183 (women)  | ≥ 70   | 54           | 85 (76-91)             | 86 (76-92)             | 88 (81-94)     | 83 (75-91)     | 5.94 (3.49-10.09)   | 0.18 (0.11-0.28) |
| Merkle, et al., 2010 <sup>96</sup>       | Mixed        | Good    | 77 (women)   | ≥ 50   | 56           | 91 (78-97)             | 91 (76-98)             | 93 (85-100)    | 89 (78-99)     | 10.28 (3.47-30.4)   | 0.10 (0.04-0.26) |
|  |              |         | 179 (men)    |        | 79           | 91 (86-96)             | 74 (57-87)             | 93 (89-97)     | 70 (56-84)     | 3.48 (2.04-5.93)    | 0.12 (0.06-0.20) |
| Coelho-Filho, et al., 2011 <sup>99</sup> | Mixed        | Fair    | 168 (women)  | ≥ 70   | 7            | 92 (62-100)            | 80 (44-97)             | 85 (65-100)    | 89 (68-100)    | 4.58 (1.31-16.02)   | 0.10 (0.01-0.70) |
|  |              |         | 237 (men)    |        | 15           | 86 (71-95)             | 74 (49-91)             | 86 (75-97)     | 74 (54-93)     | 3.27 (1.52-7.02)    | 0.19 (0.08-0.44) |

Abbreviations: CAD = coronary artery disease; Cath % = % stenosis defined to be positive for CAD on diagnostic cardiac catheterization (coronary angiography); CI = confidence interval; LR+ = positive likelihood ratio; LR- = negative likelihood ratio; NPV = negative predictive value; NR = not reported; PPV = positive predictive value



## Coronary CTA

We identified eight studies evaluating the accuracy of coronary CTA compared with coronary angiography (Table 6).<sup>66,98,100-105</sup> Five of these studies were exclusively in women with no known CAD, and these are the studies used in our primary analysis. In our secondary analysis, we evaluated the accuracy of coronary CTA in diagnosing CAD in mixed populations of known and no known CAD, including the three additional studies.

### Primary Analysis: Population of Women With No Known CAD

The five studies represent findings on coronary CTA use in 474 women (sample size ranging from 30 to 280 women). Of these studies, three were good quality, two were fair quality, and none were poor quality. Sensitivity varied from 70 to 100 percent, and specificity varied from 46 to 91 percent; the median sensitivity was 89 percent, and the median specificity was 78 percent. Figure 34 presents forest plots of the individual study estimates of sensitivity and specificity of coronary CTA for diagnosing CAD in women with no known CAD.

**Figure 34. Accuracy of coronary CTA in women with no known CAD**

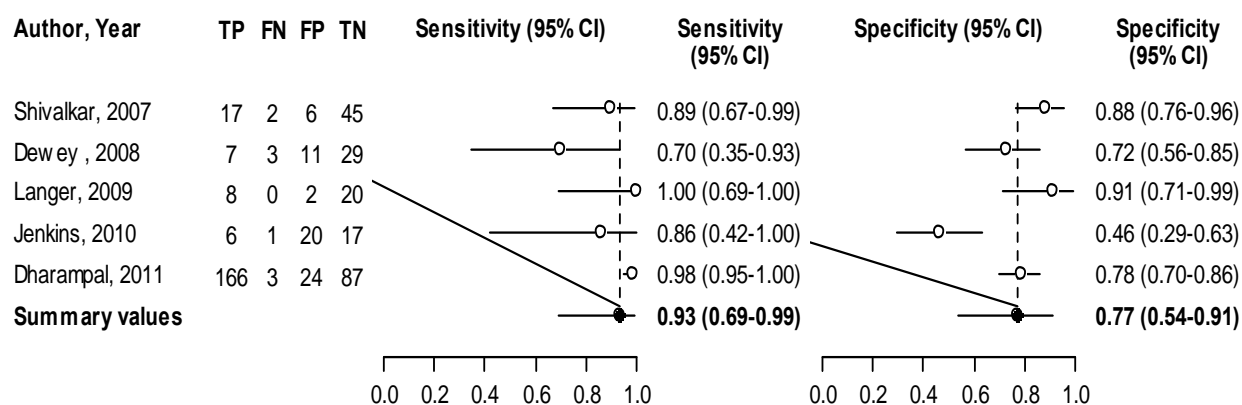
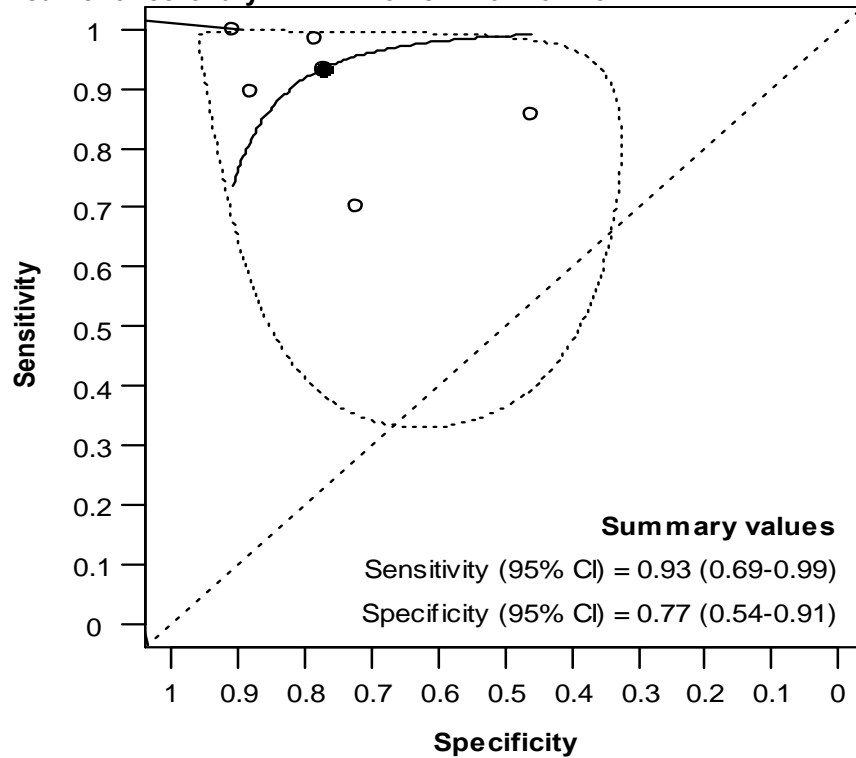


Figure 35 presents an SROC curve with an average sensitivity of 93 percent (95% CI, 69 to 99 percent) and specificity of 77 percent (95% CI, 54 to 91 percent).

**Figure 35. SROC curve for coronary CTA in women with no known CAD**



The prevalence of CAD in the 5 studies ranged from 16 to 60 percent with a mean prevalence of 30 percent. In the individual studies, PPV ranged from 23 to 87 percent, and NPV ranged from 46 to 91 percent. LR+ ranged from 1.58 to 11 and LR- from 0 to 0.41. Using the summary sensitivity and specificity of 93 and 77 percent, respectively, we calculated an overall PPV of 63 percent and NPV of 96 percent. Similarly, we calculated summary LR+ of 4.04 and LR- of 0.09.

### **Accuracy of Coronary CTA in Three Good-Quality Studies**

Next, we evaluated the accuracy of coronary CTA compared with coronary angiography in the three good-quality studies. In these studies, sensitivity varied from 70 to 100 percent, and specificity varied from 46 to 91 percent; the median sensitivity was 86 percent, and the median specificity was 73 percent. Given the small number of studies and the specific point estimate and CIs of these studies, our meta-analytic modeling was not able to reach convergence and provide a summary sensitivity and specificity for this set of studies.

The prevalence of CAD in these 3 good-quality studies ranged from 16 to 27 percent with a mean prevalence of 21 percent. In the individual studies, PPV ranged from 23 to 80 percent, and NPV ranged from 91 to 100 percent. LR+ ranged from 1.58 to 11 and LR- from 0 to 0.41.

## Secondary Analysis: Mixed Population of Women With Known and No Known CAD

We performed a secondary analysis where we expanded our inclusion criteria to include studies whose patient population included a mix of women with known CAD and women with no known CAD. This expanded inclusion criteria allowed three additional studies to be included in our analysis (totaling eight studies). The 8 studies represent findings on coronary CTA use in 690 women (sample size ranging from 30 to 280 women). Of these eight studies, four were good quality, four were fair quality, and none were poor quality (Table 6).

In these 8 studies, sensitivity varied from 70 to 100 percent, and specificity varied from 46 to 100 percent; the median sensitivity was 92 percent, and the median specificity was 88 percent. Figure 36 presents forest plots of the individual study estimates of sensitivity and specificity of coronary CTA for diagnosing CAD in women from mixed populations.

**Figure 36. Accuracy of coronary CTA in women from mixed populations**

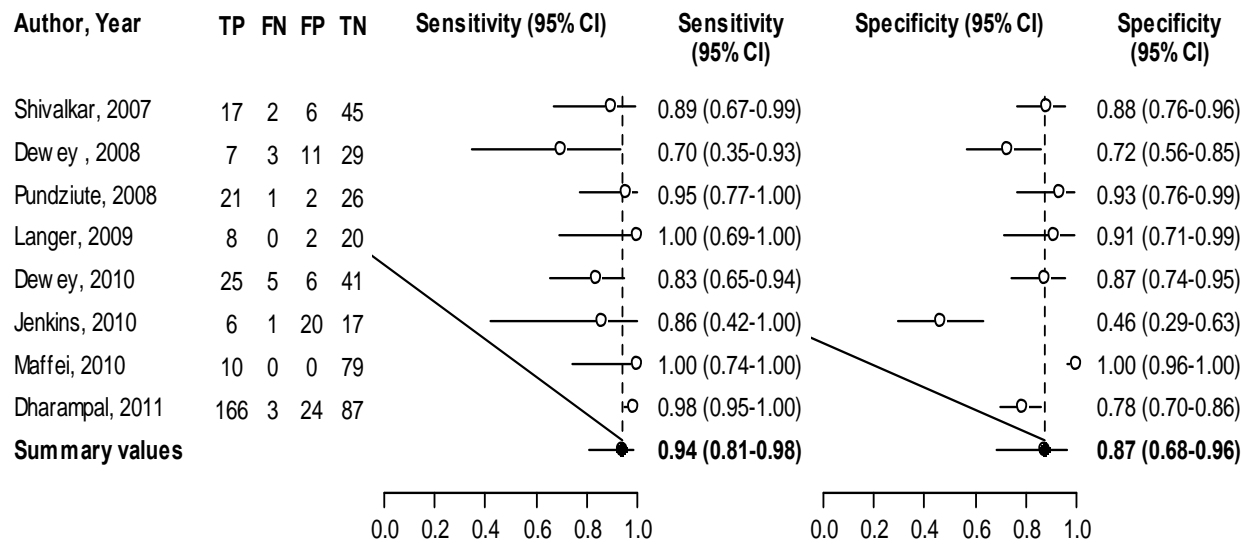
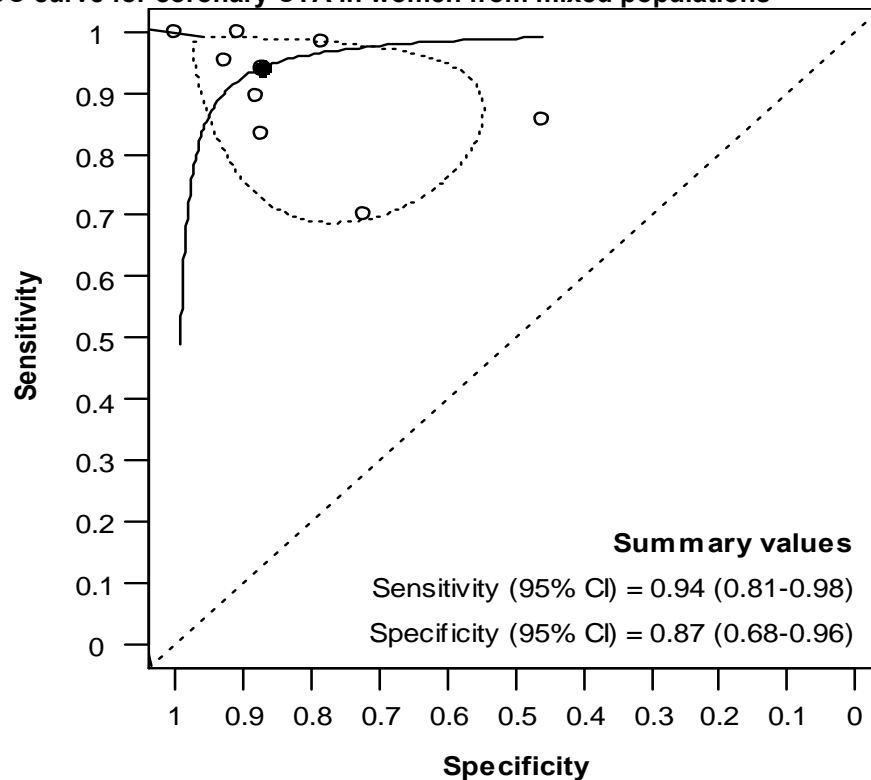


Figure 37 presents an SROC curve demonstrating an average sensitivity of 94 percent (95% CI, 81 to 98 percent) and specificity of 87 percent (95% CI, 68 to 96 percent).

Figure 37. SROC curve for coronary CTA in women from mixed populations



The prevalence of CAD in these 8 studies ranged from 11 to 60 percent with a mean prevalence of 31 percent. In the individual studies, PPV ranged from 23 to 100 percent, and NPV ranged from 46 to 100 percent. LR+ ranged from 2.54 to 13.36 and LR- 0 to 0.41. Using the summary sensitivity and specificity of 94 and 87 percent, respectively, we calculated an overall PPV of 76 percent and NPV of 97 percent. Similarly, we calculated summary LR+ of 7.23 and LR- of 0.069.

### Accuracy of Coronary CTA in Four Good-Quality Studies

Next, we evaluated the accuracy of coronary CTA compared with coronary angiography in the four good-quality studies. In these studies, sensitivity varied from 70 to 100 percent, and specificity varied from 46 to 91 percent; the median sensitivity was 85 percent, and the median specificity was 80 percent. Figure 38 presents forest plots of the individual study estimates of sensitivity and specificity of coronary CTA for diagnosing CAD in women with from mixed populations.

**Figure 38. Accuracy of coronary CTA in four good-quality studies in women from mixed populations**

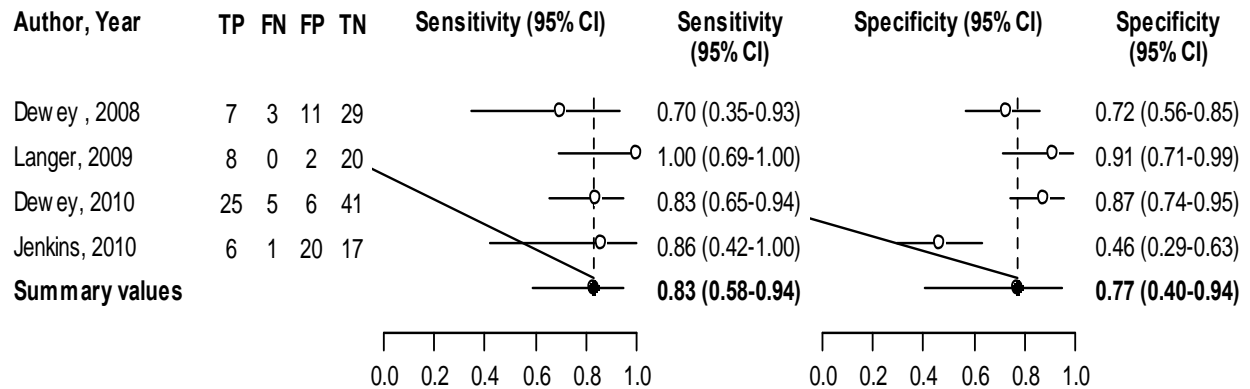
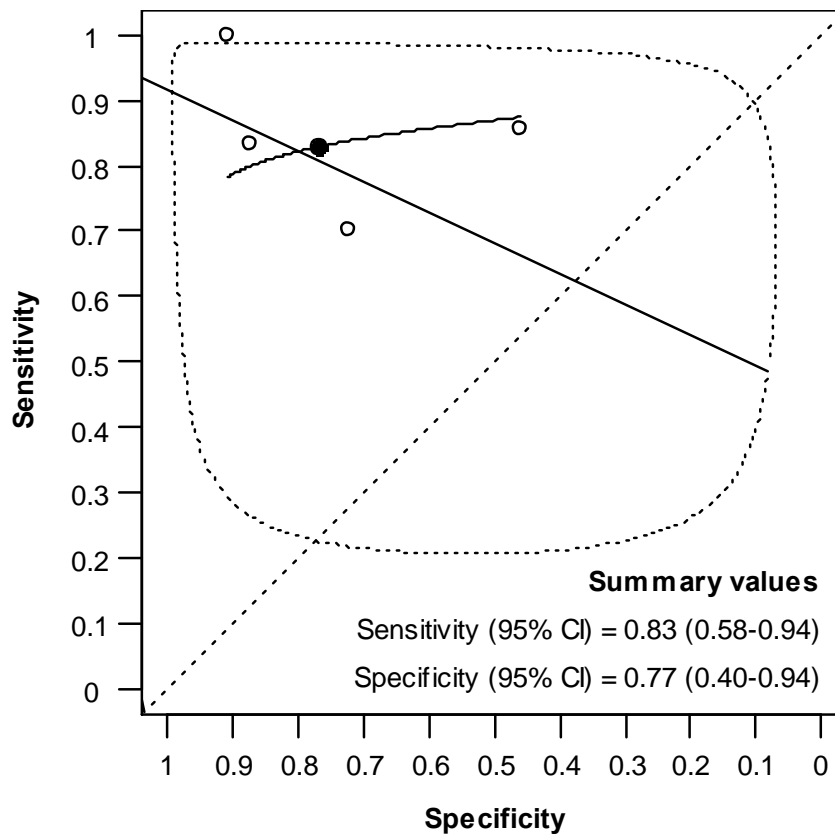


Figure 39 presents an SROC curve demonstrating an average sensitivity of 83 percent (95% CI, 58 to 94 percent) and specificity of 77 percent (95% CI, 40 to 94 percent). It is important to note that given the small number of studies and wide confidence intervals that these summary statistics should be interpreted with caution.

**Figure 39. SROC curve for coronary CTA in four good-quality studies in women from mixed populations**



The prevalence of CAD in these 4 good-quality studies ranged from 16 to 39 percent with a mean prevalence of 25 percent. In the individual studies, PPV ranged from 23 to 81 percent, and NPV ranged from 89 to 100 percent. LR+ ranged from 1.58 to 11 and LR- from 0 to 0.41. Using the summary sensitivity and specificity of 83 and 77 percent, respectively, we calculated an overall PPV of 55 percent and NPV of 93 percent. Similarly, we calculated summary LR+ of 3.61 and LR- of 0.22.

**Table 6. Summary of accuracy data evaluating coronary CTA for diagnosing CAD**

| Study                                  | Patient Mix  | Quality | Patients (N) | Cath % | Prevalence % | Sensitivity % (95% CI) | Specificity % (95% CI) | PPV % (95% CI) | NPV % (95% CI) | LR+ (95% CI)      | LR- (95% CI)       |
|--|--------------|---------|--------------|--------|--------------|------------------------|------------------------|----------------|----------------|-------------------|--------------------|
| Jenkins, et al., 2010 <sup>102</sup>   | No known CAD | Good    | 44 (women)   | ≥ 50   | 16           | 86 (42-100)            | 46 (29-63)             | 23 (7-39)      | 94 (84-100)    | 1.59 (1.04-2.42)  | 0.31 (0.05-1.97)   |
|  |              |         | 55 (men)     |        | 56           | 94 (79-99)             | 50 (29-71)             | 71 (57-85)     | 86 (67-100)    | 1.87 (1.24-2.82)  | 0.13 (0.03-0.52)   |
| Langer, et al., 2009 <sup>98</sup>     | No known CAD | Good    | 30 (women)   | ≥ 50   | 27           | 100 (69-100)           | 91 (71-99)             | 80 (55-100)    | 100 (85-100)   | 11 (2.93-41.24)   | 0 (NA)             |
|  |              |         | 38 (men)     |        | 47           | 94 (73-100)            | 100 (86-100)           | 100 (82-100)   | 95 (86-100)    | NA                | 0.06 (0.01-0.37)   |
| Dewey, et al., 2008 <sup>103</sup>     | No known CAD | Good    | 50 (women)   | ≥ 50   | 20           | 70 (35-93)             | 73 (56-85)             | 39 (16-61)     | 91 (81-100)    | 2.55 (1.33-4.86)  | 0.41 (0.16-1.09)   |
|  |              |         | 95 (men)     |        | 62           | 95 (86-99)             | 81 (64-92)             | 89 (81-97)     | 91 (81-100)    | 4.88 (2.50-9.52)  | 0.06 (0.02-0.19)   |
| Shivalkar, et al., 2007 <sup>104</sup> | No known CAD | Fair    | 70 (women)   | ≥ 70   | 27           | 89 (67-99)             | 88 (76-96)             | 74 (56-92)     | 96 (90-100)    | 7.61 (3.53-16.38) | 0.12 (0.03-0.44)   |
| Dharampal, et al., 2011 <sup>105</sup> | No known CAD | Fair    | 280 (women)  | ≥ 50   | 60           | 98 (95-100)            | 78 (70-86)             | 87 (83-92)     | 97 (93-100)    | 4.54 (3.19-6.48)  | 0.02 (0.01-0.07)   |
|  |              |         | 636 (men)    |        | 76           | 99 (98-100)            | 82 (75-88)             | 95 (93-97)     | 98 (95-100)    | 5.56 (3.95-7.82)  | 0.007 (0.002-0.02) |
| Dewey, et al., 2010 <sup>100</sup>     | Mixed        | Good    | 77 (women)   | ≥ 50   | 39           | 83 (65-94)             | 87 (74-95)             | 81 (67-95)     | 89 (80-98)     | 6.53 (3.04-14.02) | 0.19 (0.09-0.43)   |
|  |              |         | 214 (men)    |        | 62           | 86 (79-91)             | 91 (83-96)             | 94 (90-98)     | 80 (71-88)     | 9.92 (4.87-20.20) | 0.16 (0.10-0.24)   |

**Table 6. Summary of accuracy data evaluating coronary CTA for diagnosing CAD (continued)**

| Study                                  | Patient Mix | Quality | Patients (N) | Cath % | Prevalence % | Sensitivity % (95% CI) | Specificity % (95% CI) | PPV % (95% CI) | NPV % (95% CI) | LR+ (95% CI)       | LR- (95% CI)     |
|--|-------------|---------|--------------|--------|--------------|------------------------|------------------------|----------------|----------------|--------------------|------------------|
| Dewey, et al., 2010 <sup>100</sup>     | Mixed       | Good    | 77 (women)   | ≥ 50   | 39           | 83 (65-94)             | 87 (74-95)             | 81 (67-95)     | 89 (80-98)     | 6.53 (3.04-14.02)  | 0.19 (0.09-0.43) |
|  |             |         | 214 (men)    |        | 62           | 86 (79-91)             | 91 (83-96)             | 94 (90-98)     | 80 (71-88)     | 9.92 (4.87-20.20)  | 0.16 (0.10-0.24) |
| Pundziute, et al., 2008 <sup>101</sup> | Mixed       | Fair    | 50 (women)   | ≥ 50   | 44           | 95 (77-100)            | 93 (76-99)             | 91 (80-100)    | 96 (89-100)    | 13.36 (3.50-50.97) | 0.05 (0.01-0.33) |
|  |             |         | 50 (men)     |        | 64           | 100 (91-100)           | 89 (65-99)             | 94 (86-100)    | 100 (81-100)   | 9 (2.44-33.24)     | 0 (NA)           |
| Maffei, et al., 2010 <sup>66</sup>     | Mixed       | Fair    | 89 (women)   | ≥ 50   | 11           | 100 (74-100)           | 100 (96-100)           | 100 (70-100)   | 100 (96-100)   | NA                 | 0 (NA)           |
|  |             |         | 88 (men)     |        | 18           | 100 (83-100)           | 97 (90-100)            | 89 (74-100)    | 100 (96-100)   | 36 (9.18-141)      | 0 (NA)           |

Abbreviations: CAD = coronary artery disease; Cath % = % stenosis defined to be positive for CAD on diagnostic cardiac catheterization (coronary angiography); CI = confidence interval; LR+ = positive likelihood ratio; LR- = negative likelihood ratio; NPV = negative predictive value; NR = not reported; PPV = positive predictive value



## KQ 1 Summary

Table 7 and Figure 40 show the summary of the diagnostic accuracy of ECG, ECHO, SPECT, CMR, and coronary CTA modalities in women presented in Figures 4–39. The information is presented separately for no known CAD and mixed CAD populations, as well as for all studies separately from the good-quality studies. Overall, within a given modality, the summary sensitivities and specificities were similar for both types of populations (known and no known CAD) and for all studies when compared with good-quality studies. When accounting for only the good-quality studies, it appeared that the diagnostic accuracy of detecting CAD in women with unknown CAD was better (in descending order) for coronary CTA, SPECT, ECHO, CMR, and ECG. For the newer technologies (i.e., coronary CTA and CMR), more studies in women would be needed to support these findings since the 95% CIs were quite wide.

To minimize the risk of spectrum bias, our primary analysis focused on women with no known CAD. We also explored mixed populations of women with known and no known CAD in sensitivity analyses. These analyses did not demonstrate a significant difference in terms of the sensitivities and specificities from our primary analysis. We also explored whether the accuracy of the modalities were correlated with the underlying prevalence of disease in the population of interest. The mean prevalences and 95% CIs for ECG, SPECT, ECHO, CMR, and coronary CTA with the population of women with no previously known CAD were 0.41 (0.36 to 0.46), 0.44 (0.34 to 0.55), 0.43 (0.37 to 0.50), 0.26 (0.14 to 0.44), and 0.29 (0.13 to 0.54), respectively. We evaluated whether these prevalences were different across modalities using a random-effects model and did not find a statistically significant difference ( $p = 0.17$ ). Thus, these analyses did not indicate any specific trend or relationship between prevalence and the NIT's sensitivity or specificity. There did appear to be an increase in the sensitivity of CMR over time, although the wide confidence intervals for this characteristic highlight the uncertainty in this trend.

We assessed the risk of verification bias by exploring the studies in our analysis that did not complete a coronary catheterization in all of the patients who underwent the NIT. In the population of women with no previously known CAD, this represented one study of SPECT,<sup>52</sup> one study of ECHO,<sup>79</sup> three studies of ECG,<sup>29,52,58</sup> and no studies of CMR or coronary CTA. Given the small number of total studies with this potential bias, we felt confident that our primary results were minimized for verification bias.

We explored the potential for publication bias across the different modalities in our four populations of interest (studies of women with no known CAD, good-quality studies of women with no known CAD, studies of women from mixed populations, and good-quality studies of women from mixed populations). Our analyses did not provide evidence for publication bias, with our  $p$  values ranging from 0.093 to 0.95.

In a final analysis, we explored whether there was a statistically significant difference between the diagnostic accuracy of testing modalities in women using a generalized linear mixed model, with NIT modality and disease state (no known and mixed CAD) as covariates in the model. Our analyses determined that for women with no previously known CAD, there were differences between the performance of the available modalities ( $p < 0.001$ ). The sensitivity of ECHO and SPECT was significantly higher than that of ECG. Specificity of ECG was less than that of CMR (borderline) and of ECHO. We similarly explored the differences among the modalities in the subset of studies that were good-quality and also where there was no known CAD in the included population. These analyses again demonstrated differences between

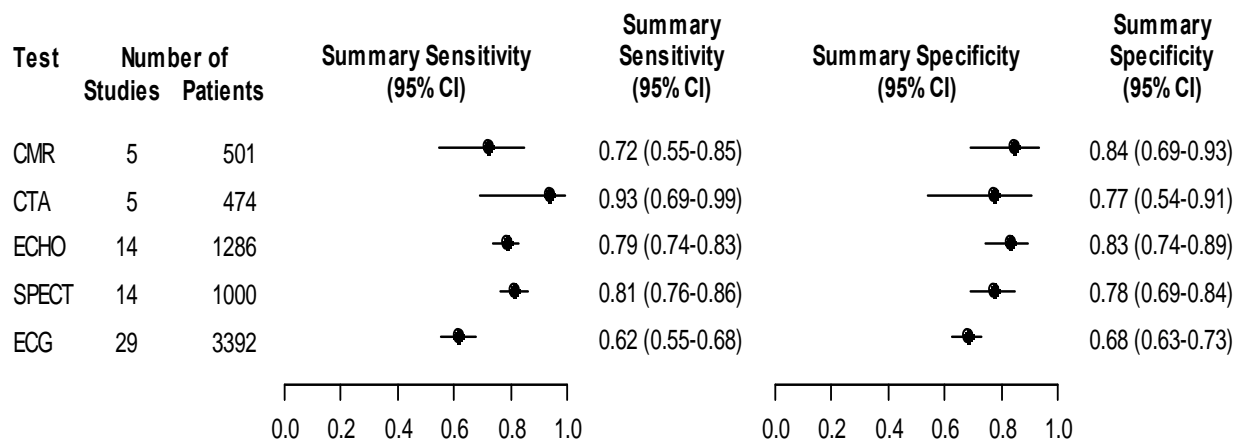
performance of tests ( $p = 0.008$ ) with the specificity of ECG being less than that of CMR and ECHO.

**Table 7. Summary of accuracy of NITs compared with coronary angiography for diagnosing CAD in women**

| Modality     | Population       | Quality of Included Studies | Number of Studies | Number of Women | Summary Sensitivity (95% CI) | Summary Specificity (95% CI) |
|--------------|------------------|-----------------------------|-------------------|-----------------|------------------------------|------------------------------|
| ECG          | No known CAD     | All                         | 29                | 3392            | 62% (55%-68%)                | 68% (63%-73%)                |
|              |                  | Good                        | 10                | 1410            | 70% (58%-79%)                | 62% (53%-69%)                |
|              | Mixed population | All                         | 41                | 4879            | 61% (54%-67%)                | 65% (58%-72%)                |
|              |                  | Good                        | 13                | 1679            | 65% (52%-76%)                | 60% (52%-68%)                |
| ECHO         | No known CAD     | All                         | 14                | 1286            | 79% (74%-83%)                | 83% (74%-89%)                |
|              |                  | Good                        | 5                 | 561             | 79% (69%-87%)                | 85% (68%-94%)                |
|              | Mixed population | All                         | 22                | 1873            | 78% (73%-83%)                | 86% (79%-91%)                |
|              |                  | Good                        | 8                 | 807             | 77% (65%-85%)                | 89% (76%-95%)                |
| SPECT        | No known CAD     | All                         | 14                | 1000            | 81% (76%-86%)                | 78% (69%-84%)                |
|              |                  | Good                        | 4                 | 394             | 83% (52%-95%)                | 72% (37%-92%)                |
|              | Mixed population | All                         | 30                | 2146            | 82% (77%-87%)                | 81% (74%-86%)                |
|              |                  | Good                        | 10                | 982             | 82% (72%-88%)                | 79% (66%-87%)                |
| CMR          | No known CAD     | All                         | 5                 | 501             | 72% (55%-85%)                | 84% (69%-93%)                |
|              |                  | Good                        | 5                 | 501             | 72% (55%-85%)                | 84% (69%-93%)                |
|              | Mixed population | All                         | 6                 | 778             | 78% (61%-89%)                | 84% (74%-90%)                |
|              |                  | Good                        | 5                 | 610             | 76% (55%-89%)                | 84% (72%-91%)                |
| Coronary CTA | No known CAD     | All                         | 5                 | 474             | 93% (69%-99%)                | 77% (54%-91%)                |
|              |                  | Good                        | 3                 | 124             | 85% (26%-99%)                | 73% (17%-97%)                |
|              | Mixed population | All                         | 8                 | 690             | 94% (81%-98%)                | 87% (68%-96%)                |
|              |                  | Good                        | 4                 | 201             | 83% (58%-94%)                | 77% (40%-94%)                |

Abbreviations: CAD = coronary artery disease; CI = confidence interval; CMR =cardiac magnetic resonance; CTA = computed tomography angiography; ECG = exercise/stress electrocardiogram; ECHO = echocardiogram; SPECT = single proton emission computed tomography

**Figure 40. Summary of accuracy of NITs compared with coronary angiography for diagnosing CAD in women with no known CAD (all studies)**



### Comparative Accuracy of NIT Modalities in Men

Although it was not the primary goal of this systematic review, we also evaluated, when possible, the accuracy of the five NIT modalities in male patients and specifically how the accuracy of these modalities differed between men and women. Most of the studies included in our analysis, however, did not include data on both sexes. Specifically, of the 41 included studies evaluating ECG in a mixed population, only 20 included data on men as well as women. Similarly for the 22 ECHO, 30 SPECT, 6 CMR, and 8 CTA included studies, only 9, 11, 3, and 7 respectively included data on men.

Although limited, the available studies provided enough data for men to determine summary sensitivity and specificity estimates and to evaluate whether the accuracy of these modalities differed between men and women (Table 8). In Tables 2–6, we provide the accuracy data for the individual studies included in our analysis that had male representation. In Table 7 we list the summary sensitivities and specificities calculated with an SROC curve as described in our primary women analyses. Given the reduced number of available studies, we focused on studies with populations of either no known CAD or a mix of known and no known CAD. When comparing the accuracy of the modalities between men and women enrolled in the same studies, the ECG and coronary CTA modalities were both less sensitive and less specific in women. The ECHO, CMR, and SPECT modalities, although less sensitive, appeared to be more specific in women. The lower specificity of the ECG modality in women, however, is the only estimate that was determined to be a statistically significant difference.

**Table 8. Summary of accuracy of NITs for diagnosing CAD in men compared with women from mixed populations**

| Modality     | Quality of Included Studies | Number of Studies | Number of Men | Summary Sensitivity in Men (95% CI) | Summary Sensitivity in Women (95% CI) | p Value (Women vs. Men) | Summary Specificity in Men (95% CI) | Summary Specificity in Women (95% CI) | p Value (Women vs. Men) |
|--------------|-----------------------------|-------------------|---------------|-------------------------------------|---------------------------------------|-------------------------|-------------------------------------|---------------------------------------|-------------------------|
| ECG          | All                         | 20                | 7345          | 64% (54%-73%)                       | 61% (54%-67%)                         | 0.57                    | 81% (72%-87%)                       | 65% (58%-72%)                         | 0.007                   |
|              | Good                        | 5                 | 2410          | 68% (32%-90%)                       | 65% (52%-76%)                         |                         | 74% (49%-90%)                       | 60% (52%-68%)                         |                         |
| ECHO         | All                         | 9                 | 1705          | 77% (65%-86%)                       | 78% (73%-83%)                         | 0.80                    | 81% (65%-91%)                       | 86% (79%-91%)                         | 0.50                    |
|              | Good                        | 2                 | 666           | Insufficient data                   | 77% (65%-85%)                         |                         | Insufficient data                   | 89% (76%-95%)                         |                         |
| SPECT        | All                         | 11                | 1433          | 88% (73%-95%)                       | 82% (77%-87%)                         | 0.36                    | 74% (50%-89%)                       | 81% (74%-86%)                         | 0.47                    |
|              | Good                        | 3                 | 323           | 91% (78%-97%)                       | 82% (72%-88%)                         |                         | 71% (32%-93%)                       | 79% (66%-87%)                         |                         |
| CMR          | All                         | 3                 | 272           | 86% (50%-97%)                       | 78% (61%-89%)                         | 0.53                    | 72% (46%-89%)                       | 84% (74%-90%)                         | 0.12                    |
|              | Good                        | 2                 | 217           | Insufficient data                   | 76% (55%-89%)                         |                         | Insufficient data                   | 84% (72%-91%)                         |                         |
| Coronary CTA | All                         | 7                 | 1176          | 97% (89%-99%)                       | 94% (81%-98%)                         | 0.36                    | 89% (71%-96%)                       | 87% (68%-96%)                         | 0.87                    |
|              | Good                        | 4                 | 402           | 91% (77%-97%)                       | 83% (58%-94%)                         |                         | 85% (43%-98%)                       | 77% (40%-94%)                         |                         |

Abbreviations: CAD = coronary artery disease; CI = confidence interval; CMR =cardiac magnetic resonance; CTA = computed tomography angiography; ECG = exercise/stress electrocardiogram; ECHO = echocardiogram; SPECT = single proton emission computed tomography

## Key Question 2: Predictors of Diagnostic Accuracy

KQ 2. What are the predictors of diagnostic accuracy (e.g., age, race/ethnicity, body size, heart size, menopausal status, functional status, stress modality) of different NITs in women?

### Key Points

- Significant variability existed around diagnostic accuracy among studies examining each NIT modality.
- Only one study compared diagnostic accuracy for women with diagnostic accuracy for men.
- The studies reviewed did not examine functional status, age, or body size—the main predictors examined included heart size, pretest probability of CAD, race/ethnicity, postmenopausal status, and beta blocker use.
- There was heterogeneity in the types of predictors reported.
- Studies that examined heart size varied on the effect on diagnostic accuracy by stress modality. These studies suggest that increased heart size reduces the specificity of stress ECG, ECHO, and SPECT.
- One fair-quality study of 51 women reported that beta blocker use reduces the specificity of stress ECG and the sensitivity and specificity of SPECT. Withholding beta blockers prior to exercise stress testing is common to allow patients to achieve a target (or higher) heart rate in assessing for ischemia. One study showed that the PPV increases as the pretest probability of CAD increases for stress ECG and ECHO.
- Insufficient evidence was available to draw definitive conclusions about predictors given the small number of studies for each predictor and for each modality, as well as the combination of predictor by modality.

### Detailed Synthesis

Many factors are reported in the literature that affect the diagnostic accuracy of noninvasive testing in women, including (1) higher prevalence in women of nonobstructive CAD (microvascular abnormalities, mitral valve prolapse), (2) less predictive symptomatology, (3) limited exercise tolerance because of older age, obesity, and diabetes at initial diagnosis, (4) different response to exercise than men, (5) lower peak exercise values, (6) lower increase in the left ventricular ejection fraction, (7) an increase in cardiac output by enhancing end-diastolic volume, (8) inappropriate catecholamine release, (9) hormonal influences of estrogens mimicking a digitalis-like false-positive ECG response, (10) anatomic differences affecting stress test results, (11) breast attenuation artifacts, (12) smaller coronary artery size, (13) smaller left ventricular chamber size, (14) higher prevalence of single-vessel disease, and (15) poor left ventricular opacification on echocardiography.

For KQ 2, we examined studies for the following nine predictors of diagnostic accuracy of different NITs in women: age, race/ethnicity, body size, heart size, menopausal status, functional status, stress modality, cardiac risk factors, and pretest probability of CAD. We identified 11 studies<sup>25,28,31,38,50,55,67,69,83,95,97</sup> that described diagnostic accuracy, but only 9 of these studies examined *predictors* of diagnostic accuracy. Four of the 11 studies were considered good quality, 5 were fair quality, and 2 were poor quality.

## Findings of Diagnostic Accuracy by Predictor

Findings of diagnostic accuracy by predictor in the studies we reviewed are summarized in Tables 9–12. Of the nine predictors we originally searched for, we found studies that addressed four of the predictors: (1) age combined with (2) menopausal status, (3) race/ethnicity, and (4) heart size. We also found two other types of predictors reported in women: (5) pretest probability based on cardiac risk factors and (6) use of beta blockers:

- The study by Cin,, et al.<sup>31</sup> examined the diagnostic accuracy of stress ECG in postmenopausal women ages 55 to 64.
- Two studies examined race/ethnicity: One study by Vashist,, et al.<sup>83</sup> compared the diagnostic accuracy of SPECT across three race/ethnic categories—African American, Hispanic, and Asian—and another study by Yeih,, et al.<sup>50</sup> examined the diagnostic accuracy of ECG testing and SPECT in Asian women in Taiwan.
- Four studies examined heart size as a predictor of diagnostic accuracy: Lu,, et al.<sup>28</sup> and Gebker,, et al.,<sup>95</sup> compared left ventricular hypertrophy (LVH) with no LVH for ECG, ECHO, SPECT, and CMR. The study by Klem,, et al.,<sup>97</sup> examined heart size in grams for CMR testing, and Siegler,, et al.,<sup>25</sup> examined heart size for ECG alone.
- Three studies, Yeih,, et al.,<sup>50</sup> Marwick,, et al.,<sup>55</sup> and Ho,, et al.,<sup>67</sup> examined pretest probability as a predictor of accuracy for ECG, dobutamine ECHO, and CMR.
- The study by Yeih,, et al.,<sup>50</sup> examined the use of beta blockers on diagnostic accuracy in ECG and SPECT.
- We identified no studies that examined age alone, functional status, or body size as predictors of diagnostic accuracy in women.

**Table 9. Age and menopausal status as a predictor<sup>a</sup>**

| Study (predictor)  | N Women | Study Type  | Quality Score | Modality | Sensitivity % | Specificity % | PPV % | NPV % | LR+  | LR-  |
|--|---------|-------------|---------------|----------|---------------|---------------|-------|-------|------|------|
| Cin, et al., 2000 <sup>31</sup><br>(Postmenopausal, ages 55 to 64) | 110     | Prospective | Poor          | ECG      | 86            | 61            | 81    | 70    | 2.18 | 0.23 |

<sup>a</sup>This study did not have a premenopausal reference group.

Abbreviations: ECG = exercise/stress electrocardiogram; LR+ = positive likelihood ratio; LR- = negative likelihood ratio; N = number; NPV = negative predictive value; PPV = positive predictive value

**Table 10. Race/ethnicity as a predictor<sup>a</sup>**

| Study (predictor)   | N Women | Study Type    | Quality Score | Modality | Sensitivity % | Specificity % | PPV % | NPV % | LR+  | LR-  |
|---|---------|---------------|---------------|----------|---------------|---------------|-------|-------|------|------|
| Vashist, et al., 2007 <sup>83</sup><br>(Hispanic)         | 16      | Retrospective | Fair          | SPECT    | 71.4          | 33.3          | 45.5  | 60    | 1.07 | 0.86 |
| Vashist, et al., 2007 <sup>83</sup><br>(African American) | 34      | Retrospective | Fair          | SPECT    | 90            | 20            | 86.7  | 25    | 1.12 | 0.52 |
| Yeih, et al., 2007 <sup>50</sup><br>(Asian)               | 51      | Prospective   | Fair          | SPECT    | 71            | 87            | 87    | 71    | 5.48 | 0.33 |
| Yeih, et al., 2007 <sup>50</sup><br>(Asian)               | 51      | Prospective   | Fair          | ECG      | 43            | 83            | 75    | 54    | 2.46 | 0.69 |

<sup>a</sup>None of these studies reported results for Caucasian women.

Abbreviations: ECG = exercise/stress electrocardiogram; LR+ = positive likelihood ratio; LR- = negative likelihood ratio; N = number; NPV = negative predictive value; PPV = positive predictive value; SPECT = single proton emission computed tomography

**Table 11. Heart size as a predictor**

| Study (predictor)                                | N Women              | Study Type  | Quality Score | Modality | Sensitivity %            | Specificity %          | PPV %                  | NPV %                  | LR+                        | LR-                        |
|--|----------------------|-------------|---------------|----------|--------------------------|------------------------|------------------------|------------------------|----------------------------|----------------------------|
| Lu, et al., 2010 <sup>28</sup> (With LVH)        | 36                   | Prospective | Good          | ECG      | NR                       | 31                     | NR                     | NR                     | NR                         | NR                         |
| Lu, et al., 2010 <sup>28</sup> (Without LVH)     | 40                   | Prospective | Good          | ECG      | NR                       | 69                     | NR                     | NR                     | NR                         | NR                         |
| Lu, et al., 2010 <sup>28</sup> (With LVH)        | 36                   | Prospective | Good          | SPECT    | NR                       | 31                     | NR                     | NR                     | NR                         | NR                         |
| Lu, et al., 2010 <sup>28</sup> (Without LVH)     | 40                   | Prospective | Good          | SPECT    | NR                       | 66                     | NR                     | NR                     | NR                         | NR                         |
| Lu, et al., 2010 <sup>28</sup> (With LVH)        | 36                   | Prospective | Good          | ECHO-DOB | NR                       | 69                     | NR                     | NR                     | NR                         | NR                         |
| Lu, et al., 2010 <sup>28</sup> (Without LVH)     | 40                   | Prospective | Good          | ECHO-DOB | NR                       | 89                     | NR                     | NR                     | NR                         | NR                         |
| Lu, et al., 2010 <sup>28</sup> (With LVH)        | 36                   | Prospective | Good          | ECHO-DIP | NR                       | 81                     | NR                     | NR                     | NR                         | NR                         |
| Lu, et al., 2010 <sup>28</sup> (Without LVH)     | 40                   | Prospective | Good          | ECHO-DIP | NR                       | 96                     | NR                     | NR                     | NR                         | NR                         |
| Gebker, et al., 2010 <sup>95</sup> (With LVH)    | 56 women<br>179 men  | Prospective | Good          | CMR      | 80 (women)<br>79 (men)   | 91 (women)<br>95 (men) | 93 (women)<br>98 (men) | 73 (women)<br>56 (men) | 8.4 (women)<br>15.46 (men) | 0.22 (women)<br>0.22 (men) |
| Gebker, et al., 2010 <sup>95</sup> (Without LVH) | 127 women<br>311 men | Prospective | Good          | CMR      | 87.5 (women)<br>90 (men) | 84 (women)<br>78 (men) | 85 (women)<br>91 (men) | 87 (women)<br>75 (men) | 5.51 (women)<br>4.08 (men) | 0.15 (women)<br>0.12 (men) |



**Table 11. Heart size as a predictor (continued)**

| Study (predictor)  | N Women | Study Type  | Quality Score | Modality | Sensitivity % | Specificity % | PPV % | NPV % | LR+ | LR- |
|--|---------|-------------|---------------|----------|---------------|---------------|-------|-------|-----|-----|
| Klem, et al., 2008 <sup>97</sup><br>(≤ 97 g LV mass)       | NR      | Prospective | Good          | CMR      | 69            | NR            | NR    | NR    | NR  | NR  |
| Klem, et al., 2008 <sup>97</sup><br>(≥ 97 g LV mass)       | NR      | Prospective | Good          | CMR      | 95            | NR            | NR    | NR    | NR  | NR  |
| Siegler, et al., 2011 <sup>25</sup><br>(Small heart size)  | 123     | Prospective | Fair          | ECG      | 57            | 69            | 36    | 85    | 1.9 | 0.6 |
| Siegler, et al., <sup>25</sup> 2011<br>(Normal heart size) | 359     | Prospective | Fair          | ECG      | 61            | 84            | 62    | 83    | 3.7 | 0.5 |

Abbreviations: CMR =cardiac magnetic resonance; ECG = exercise/stress electrocardiogram; ECHO-DOB = echocardiogram with dobutamine; ECHO-DIP = echocardiogram with dipyridamole; LR+ = positive likelihood ratio; LR- = negative likelihood ratio; LVH = left ventricular hypertrophy; N = number; NPV = negative predictive value; NR = not reported; PPV = positive predictive value; SPECT = single proton emission computed tomography

**Table 12. Other potential predictors**

| Study (predictor)   | N Women | Study Type  | Quality Score | Modality | Sensitivity % | Specificity % | PPV % | NPV % | LR+  | LR-   |
|---|---------|-------------|---------------|----------|---------------|---------------|-------|-------|------|-------|
| Yeih, et al., 2007 <sup>50</sup><br>(High pretest probability)            | 32      | Prospective | Fair          | ECG      | 50            | 80            | 85    | 42    | 2.5  | 0.625 |
| Yeih, et al., 2007 <sup>50</sup><br>(Low pretest probability)             | 19      | Prospective | Fair          | ECG      | 33            | 69            | 33    | 69    | 1.08 | 0.96  |
| Yeih, et al., 2007 <sup>50</sup><br>(With beta blocker)                   | 24      | Prospective | Fair          | ECG      | 53            | 56            | 67    | 42    | 1.2  | 0.84  |
| Yeih, et al., 2007 <sup>50</sup><br>(Without beta blocker)                | 27      | Prospective | Fair          | ECG      | 31            | 86            | 67    | 57    | 2.15 | 0.81  |
| Yeih, et al., 2007 <sup>50</sup><br>(With beta blocker)                   | 24      | Prospective | Fair          | SPECT    | 67            | 78            | 83    | 58    | 3.0  | 0.43  |
| Yeih, et al., 2007 <sup>50</sup><br>(Without beta blocker)                | 27      | Prospective | Fair          | SPECT    | 77            | 93            | 91    | 81    | 10.8 | 0.25  |
| Marwick, et al., 1995 <sup>55</sup><br>(High pretest probability)         | 25      | Prospective | Fair          | ECG      | 88            | 44            | 73    | 67    | 1.57 | 0.28  |
| Marwick, et al., 1995 <sup>55</sup><br>(Intermediate pretest probability) | 59      | Prospective | Fair          | ECG      | 68            | 45            | 53    | 61    | 1.24 | 0.71  |
| Marwick, et al., 1995 <sup>55</sup><br>(Low pretest probability)          | 34      | Prospective | Fair          | ECG      | 100           | 70            | 31    | 100   | 3.33 | 0     |
| Marwick, et al., 1995 <sup>55</sup><br>(High pretest probability)         | 32      | Prospective | Fair          | ECHO     | 82            | 80            | 90    | 67    | 4.09 | 0.23  |
| Marwick, et al., 1995 <sup>55</sup><br>(Intermediate pretest probability) | 72      | Prospective | Fair          | ECHO     | 76            | 86            | 78.5  | 84    | 5.44 | 0.28  |
| Marwick, et al., 1995 <sup>55</sup><br>(Low pretest probability)          | 57      | Prospective | Fair          | ECHO     | 88            | 78            | 39    | 97    | 3.90 | 0.16  |
| Ho, et al., 1998 <sup>67</sup><br>(≥2 cardiac risk factors)               | 18      | Prospective | Fair          | ECHO     | 92            | 67            | 85    | 80    | 2.75 | 0.13  |
| Ho, et al., 1998 <sup>67</sup><br>(≤1 cardiac risk factors)               | 33      | Prospective | Fair          | ECHO     | 94            | 88            | 93    | 89    | 7.53 | 0.07  |
| Ho, et al., 1998 <sup>67</sup><br>(>50% pretest probability of CAD)       | 26      | Prospective | Fair          | ECHO     | 94            | 75            | 90    | 86    | 3.78 | 0.07  |
| Ho, et al., 1998 <sup>67</sup><br>(<50% pretest probability of CAD)       | 25      | Prospective | Fair          | ECHO     | 91            | 86            | 83    | 92    | 6.36 | 0.11  |

Abbreviations: ECG = exercise/stress electrocardiogram; ECHO = echocardiogram; LR+ = positive likelihood ratio; LR- = negative likelihood ratio; N = number; NPV = negative predictive value; PPV = positive predictive value; SPECT = single proton emission computed tomography

The following sections focus on differences in predictors, organized by NIT modality. Summaries are in Tables 13–16.

## ECG

Six studies<sup>25,28,31,38,50,55</sup> of stress ECG assessed predictors of diagnostic accuracy in women, including age/menopausal status, disease probability, use of beta blockers, and heart size. There was some variability in diagnostic accuracy using ECG testing (both exercise and with dobutamine) (Table 13).

**Overall accuracy.** For exercise ECG testing, overall sensitivity ranged from 67 to 86 percent, and specificity ranged from 44 to 78 percent. Yeih,, et al.,<sup>50</sup> examined ECG testing with dobutamine, with a resulting sensitivity of 43 percent and specificity of 83 percent. Ho,, et al.,<sup>67</sup> examined ECG with exercise treadmill test and found that the sensitivity was 71 percent and specificity was 44 percent.

**Age/menopausal status.** Cin,, et al.,<sup>31</sup> examined age and menopausal status among women ages 55 to 64 (postmenopausal) and found that ECG testing performed moderately better than in other studies not targeting this age group (sensitivity of 86 percent, specificity of 60 percent). Unfortunately, that study did not have a younger or older age group to compare findings with and was rated poor quality.

**Disease probability.** Marwick,, et al.,<sup>55</sup> compared ECG testing performance across disease probability categories (low, intermediate, and high as determined by a combination of type of chest pain and age) and found that ECG testing performed better in the low-probability group compared with the intermediate- or high-probability group. The quality of this study was fair.

**Beta blockers.** In the study by Yeih,, et al.,<sup>50</sup> which compared low and high probability calculated according to age and symptoms, ECG testing performed better in the high-probability group, as expected. This study evaluated only Asian women in Taiwan. Women receiving beta blockers had fewer false positives compared with women not taking beta blockers at the time of the ECG testing.

**Heart size.** In the study by Lewis,, et al.,<sup>38</sup> ECG testing performed better after excluding nondiagnostic cases (sensitivity 67 percent versus 43 percent, specificity 78 percent versus 66 percent). In the study by Lu,, et al.,<sup>28</sup> ECG testing performed better in patients without LVH compared with those with LVH (specificity 69 percent versus 31 percent), and this finding may relate to heart size or baseline ECG strain associated with LVH.

Siegler,, et al.,<sup>25</sup> examined 1011 patients to determine if heart size could lead to higher false-positives rates among patients undergoing stress ECG; 482 women were enrolled in the study. The prevalence of CAD was 28 percent among women enrolled. Overall sensitivity and specificity for women undergoing stress ECG was 60 and 80 percent respectively. There was a significant association between ECG outcome and heart size among women ( $p = 0.03$ ), where smaller hearts were associated with a lower specificity compared with normal size hearts.

## ECHO

Four studies<sup>28,55,67,69</sup> examined the predictors of diagnostic accuracy for exercise ECHO in women, specifically disease probability and heart size (Table 14). There was significant variability in the diagnostic accuracy of stress ECHO imaging.

**Overall accuracy.** The overall sensitivity ranged from 61 to 93 percent. The specificity ranged from 71 to 91 percent. Shin,, et al.,<sup>69</sup> examined 464 patients to determine potential factors that could lead to a higher false-positive rate among patients undergoing exercise ECHO. There

were 162 women enrolled in the study. The prevalence of CAD was 34 percent among the women enrolled. For women undergoing exercise ECHO, the overall sensitivity was 82 percent and specificity was 71 percent, which was less than for men in the study. PPV was 59 percent and NPV was 88 percent; LR+ was 2.82 and LR- was 0.26.

Marwick,, et al.,<sup>55</sup> found an overall sensitivity of 80 percent ( $p = 0.050$ ) and specificity of 81 percent ( $p < 0.004$ ) for exercise ECHO compared with exercise ECG. PPV was 71 percent and NPV was 87 percent. Exercise ECHO was compared with exercise ECG, which had a sensitivity of 77 percent and specificity of 56 percent; LR+ was 4.28 and LR- was 0.25.

Lu., et al.,<sup>28</sup> examined the diagnostic accuracy of dipyridamole and dobutamine stress ECHO modalities. The study enrolled 76 Asian women from Taiwan. The prevalence of CAD was 41 percent. For dipyridamole, sensitivity was 61 percent and specificity was 91 percent. PPV was 83 percent and NPV was 77 percent; LR+ was 6.9 and LR- was 0.42. A dobutamine stress ECHO was more sensitive (87 percent) and less specific (82 percent). (Note that the p-value for comparing the sensitivity was 0.02, and for specificity the exact p-value was not reported but was said to be not statistically significant.) PPV was 77 percent and NPV was 90 percent; LR+ was 4.9 and LR- was 0.16.

**Disease probability.** The study by Marwick ,, et al.,<sup>55</sup> also compared the diagnostic accuracy of exercise ECHO in women with different pretest probabilities of CAD (i.e., high probability, intermediate probability, and low probability). The prevalence in each group was 69, 40, and 14 percent respectively. The sensitivity was highest (88 percent) among patients with low probabilities of CAD, second highest (82 percent) among those with high probability of CAD, and lowest (76 percent) among those with an intermediate probability of CAD. However, specificity was highest (86 percent) among patients with intermediate probability of CAD, the next highest (80 percent) was among patients with highest probability of CAD, and the lowest (78 percent) was among those with a low probability of CAD. The quality of this study was fair. In this study, the diagnostic accuracy changed depending on the prevalence of CAD. Ho, et al.,<sup>67</sup> examined the diagnostic accuracy of dobutamine stress ECHO. The study enrolled 51 women from Taiwan. The prevalence of CAD was 27 percent. The sensitivity was 93 percent and specificity was 82 percent; PPV was 87 percent and NPV was 90 percent; LR+ was 5.12 and LR- was 0.08. This study also compared the diagnostic accuracy of dobutamine stress ECHO in patients with different coronary risk factors and different pretest probabilities of CAD.

In women with two or more CAD risk factors, Ho, et al., found that the sensitivity was similar to the overall sensitivity (92 percent), but the specificity decreased (67 percent). PPV was 85 percent and NPV was 80 percent; LR+ was 2.75 and LR- was 0.13. For women with zero or one CAD risk factor, sensitivity was 94 percent, specificity was 88 percent; PPV was 93 percent and NPV was 89 percent; LR+ was 7.53 and LR- was 0.07.

In addition, in women with at least a 50-percent pretest probability of CAD, the sensitivity did not change dramatically from the overall sensitivity (94 percent), but the specificity increased (88 percent). PPV was 90 percent and NPV was 86 percent; LR+ was 3.78 and LR- was 0.07. In women with a less than 50 percent pretest probability of CAD, the sensitivity was 91 percent and specificity was 86 percent. PPV was 83 percent and NPV was 92 percent; LR+ was 6.36 and LR- was 0.11. One limitation of this study is that neither beta blockers nor calcium channel blockers were withheld.

**Heart size.** Heart size was also examined in the study by Lu, et al.,<sup>28</sup> specifically, the effect that LVH had on diagnostic accuracy. Studies have suggested that LVH is commonly associated with ECG repolarization abnormalities, such as ST elevations in the absence of wall motion

abnormalities or other evidence of inducible ischemia, leading to a potentially higher false-positive rate.<sup>106</sup> The presence of LVH led to a lower specificity in both dipyridamole and dobutamine (81 percent versus 69 percent) stress test modalities. The sensitivities were not reported. The overall quality of this study was good.

## SPECT

Four studies<sup>28,50,67,83</sup> of SPECT assessed predictors of diagnostic accuracy in women, including race/ethnicity, heart size and use of beta blockers. There was considerable variability in diagnostic accuracy using SPECT (Table 15).

**Overall accuracy.** Overall sensitivity ranged from 71 to 90 percent and specificity ranged from 27 to 88 percent. In the study by Ho, et al.,<sup>67</sup> the prevalence of CAD was 27 percent. This study enrolled 51 women, but only 44 of them received SPECT. The overall sensitivity and specificity for SPECT was 79 percent and 75 percent respectively. PPV was 79 percent and NPV was 75 percent; LR+ was 3.17 and LR- was 0.28. It should be noted that, as in the Vashist, et al., study, SPECT was performed by either dipyridamole or dobutamine infusions. Determination of which agent to be used was based on patient preference.

**Race/ethnicity.** Vashist, et al.,<sup>83</sup> examined race/ethnicity differences in performance of SPECT. This retrospective study was noted to have several limitations, including combining exercise SPECT with dipyridamole and dobutamine. This study found a difference in prevalence of CAD but did not find a difference in diagnostic accuracy between Hispanics and African Americans. In the study by Yeih, et al.,<sup>50</sup> that focused on Asian women, the sensitivity was similar to other race/ethnic groups, but the specificity was significantly higher than that observed in Vashist, et al.,<sup>83</sup> (87 versus 27 percent).

**Beta blockers.** In the study by Yeih, et al.,<sup>50</sup> beta blockers reduced the diagnostic accuracy compared with no beta blockers (sensitivity 67 versus 77 percent, specificity 78 versus 93 percent). This fits with current clinical practice where beta blockers are withheld 48 hours prior to exercise stress to allow patients to achieve a higher (or target) heart rate for the assessment of ischemia.

**Heart size.** In the study by Lu, et al.,<sup>28</sup> SPECT was found to have fewer false positives in the group without LVH versus the group with LVH (specificity 66 versus 31 percent).

## CMR

Two studies<sup>95,97</sup> of CMR examined the influence of heart size on diagnostic accuracy in women (Table 16). The overall sensitivities were similar at 84 percent versus 85 percent, and the specificities ranged from 86 to 88 percent.

**Overall accuracy.** Klem, et al.,<sup>97</sup> studied 136 women at two academic medical centers who presented with chest pain. A multicomponent cardiac MRI (CMR test) that consisted of adenosine stress, rest perfusion, and delayed-enhancement CMR was used. The overall sensitivity and specificity of all three modes were found to be 70 and 81 percent respectively. PPV was 74 percent and NPV was 85 percent. These were reflected in a patient sample where the prevalence of significant CAD was 27 percent. LR+ was 5.96 and LR- was 0.35. Sensitivity and specificity decreased to 78 and 56 percent if perfusion CMR was used alone; LR+ also decreased to 1.77, and LR- increased to 0.39. Also, when coronary angiography results were examined for each patient, the sensitivity for multicomponent CMR was highest when patients with significant CAD had at least two-vessel disease with a sensitivity of 100 percent, as opposed to 71 percent when the patients had only single-vessel disease. The specificity was

unaffected by the extent of CAD and remained 88 percent regardless if the patient had single-vessel or multiple-vessel disease. The overall quality of the study was good.

The study conducted by Gebker, et al.<sup>95</sup> examined the effectiveness of dobutamine stress CMR in the detection of CAD in women compared with men. The study evaluated 183 women and 541 men for suspected CAD. The prevalence of CAD in women and men was 54 percent (99 women) and 74 percent (365 men). The overall sensitivity and specificity of dobutamine stress CMR was found to be 85 and 86 percent in women and 86 and 83 percent in men. PPV and NPV were 88 and 83 percent in women and 94 and 67 percent in men; LR+ was 5.94 and LR- was 0.18 in women compared with 5.12 and 0.17 in men. The sensitivity of detecting CAD was greater in the presence of multiple-vessel CAD than in single-vessel CAD in both men and women (91 percent compared with 81 percent in women and 91 percent compared with 82 percent in men).

**Heart size.** Klem, et al.,<sup>97</sup> also analyzed study data to determine if the heart size of patients affected the ability to detect CAD. Investigators divided the patients into two groups—those with LV mass less than 97 grams (defined as “small” hearts) and those with LV mass greater than 97 grams (defined as “large” hearts). The prevalence of significant CAD was found to be similar and was 29 percent in those with small hearts and 26 percent in those with large hearts. The reported sensitivity was 69 percent in those with small hearts and 95 percent in those with large hearts. The authors suggested that this was caused by the limitations in spatial resolution with the 1.5 Tesla MR magnet and by the fact that a smaller heart leads to a smaller number of image pixels that are available to visualize the left ventricular wall.<sup>97</sup>

Gebker, et al.,<sup>95</sup> also attempted to understand the relationship between heart size, in particular LVH, and the ability to detect the presence of CAD in women compared with men. Previous studies have shown that the presence of LVH causes a higher rate of false-negative studies.<sup>107</sup> Women undergoing dobutamine CMR with evidence of LVH had a lower sensitivity but higher specificity than women without LVH (sensitivity 80 percent versus 87.5 percent, specificity 91 percent versus 84 percent). Men undergoing dobutamine CMR with evidence of LVH had a lower sensitivity and a higher specificity than men without LVH (sensitivity 79 percent versus 90 percent, specificity 95 percent versus 78 percent).

**Table 13. Summary table for ECG**

| Predictor                        | Study                               | N Women | Study Type  | Quality Score | Sensitivity % | Specificity % | PPV % | NPV % | LR+  | LR-   |
|----------------------------------|-------------------------------------|---------|-------------|---------------|---------------|---------------|-------|-------|------|-------|
| Overall (Dobutamine)             | Yeih, et al., 2007 <sup>50</sup>    | 51      | Prospective | Fair          | 43            | 83            | 75    | 54    | 2.46 | 0.69  |
| High pretest probability         | Yeih, et al., 2007 <sup>50</sup>    | 32      | Prospective | Fair          | 50            | 80            | 85    | 42    | 2.5  | 0.625 |
| Low pretest probability          | Yeih, et al., 2007 <sup>50</sup>    | 19      | Prospective | Fair          | 33            | 69            | 33    | 69    | 1.08 | 0.96  |
| With beta blocker                | Yeih, et al., 2007 <sup>50</sup>    | 24      | Prospective | Fair          | 53            | 56            | 67    | 42    | 1.2  | 0.84  |
| Without beta blocker             | Yeih, et al., 2007 <sup>50</sup>    | 27      | Prospective | Fair          | 31            | 86            | 67    | 57    | 2.15 | 0.81  |
| Asian                            | Yeih, et al., 2007 <sup>50</sup>    | 51      | Prospective | Fair          | 43            | 83            | 75    | 54    | 2.46 | 0.69  |
| Overall                          | Ho, et al., 1998 <sup>67</sup>      | 30      | Prospective | Fair          | 71            | 44            | 0.53  | 0.64  | 1.27 | 0.65  |
| Overall                          | Cin, et al., 2000 <sup>31</sup>     | 110     | Prospective | Poor          | 86            | 61            | 81    | 70    | 2.18 | 0.23  |
| Postmenopausal, ages 55 to 64    | Cin, et al., 2000 <sup>31</sup>     | 110     | Prospective | Poor          | 86            | 61            | 81    | 70    | 2.18 | 0.23  |
| Overall                          | Marwick, et al., 1995 <sup>55</sup> | 118     | Prospective | Fair          | 77            | 56            | 54    | 78    | 1.74 | 0.41  |
| High pretest probability         | Marwick, et al., 1995 <sup>55</sup> | 25      | Prospective | Fair          | 88            | 44            | 73    | 67    | 1.57 | 0.28  |
| Intermediate pretest probability | Marwick, et al., 1995 <sup>55</sup> | 59      | Prospective | Fair          | 68            | 45            | 53    | 61    | 1.24 | 0.71  |
| Low probability                  | Marwick, et al., 1995 <sup>55</sup> | 34      | Prospective | Fair          | 100           | 70            | 31    | 100   | 3.33 | 0     |
| Overall                          | Lewis, et al., 2005 <sup>38</sup>   | 96      | Prospective | Good          | 67            | 78            | 83    | 58    | 3.0  | 0.43  |
| Excluding nondiagnostic          | Lewis, et al., 2005 <sup>38</sup>   | 74      | Prospective | Good          | 43            | 66            | 33    | 74    | 1.26 | 0.86  |
| Overall                          | Lu, et al., 2010 <sup>28</sup>      | 76      | Prospective | Good          | 81            | 56            | 56    | 81    | 1.81 | 0.35  |
| With LVH                         | Lu, et al., 2010 <sup>28</sup>      | 36      | Prospective | Good          | NR            | 31            | NR    | NR    | NR   | NR    |
| Without LVH                      | Lu, et al., 2010 <sup>28</sup>      | 40      | Prospective | Good          | NR            | 69            | NR    | NR    | NR   | NR    |
| Small heart size                 | Siegler, et al., 2011 <sup>25</sup> | 123     | Prospective | Fair          | 57            | 69            | 36    | 85    | 1.9  | 0.6   |
| Normal heart size                | Siegler, et al., 2011 <sup>25</sup> | 359     | Prospective | Fair          | 61            | 84            | 62    | 83    | 3.7  | 0.5   |

Abbreviations: LR+ = positive likelihood ratio; LR- = negative likelihood ratio; LVH = left ventricular hypertrophy; CMR = cardiac magnetic resonance; N = number; NPV = negative predictive value; NR = not reported; PPV = positive predictive value

**Table 14. Summary table for ECHO**

| Predictor                        | Study                               | N Women | Study Type  | Quality Score | Sensitivity % | Specificity % | PPV % | NPV % | LR+  | LR-  |
|----------------------------------|-------------------------------------|---------|-------------|---------------|---------------|---------------|-------|-------|------|------|
| Overall                          | Shin, et al., 2003 <sup>69a</sup>   | 162     | Prospective | Fair          | 82            | 71            | 59    | 88    | 2.82 | 0.26 |
| Overall                          | Marwick, et al., 1995 <sup>55</sup> | 161     | Prospective | Fair          | 80            | 81            | 71    | 87    | 4.28 | 0.25 |
| High pretest probability         | Marwick, et al., 1995 <sup>55</sup> | 32      | Prospective | Fair          | 82            | 80            | 90    | 67    | 4.09 | 0.23 |
| Intermediate pretest probability | Marwick, et al., 1995 <sup>55</sup> | 72      | Prospective | Fair          | 76            | 86            | 78.5  | 84    | 5.44 | 0.28 |
| Low pretest probability          | Marwick, et al., 1995 <sup>55</sup> | 57      | Prospective | Fair          | 88            | 78            | 39    | 97    | 3.90 | 0.16 |
| Dipyridamole                     | Lu, et al., 2010 <sup>28</sup>      | 76      | Prospective | Good          | 61            | 91            | 83    | 77    | 6.9  | 0.42 |
| With LVH                         | Lu, et al., 2010 <sup>28</sup>      | 36      | Prospective | Good          | NR            | 81            | NR    | NR    | NR   | NR   |
| Without LVH                      | Lu, et al., 2010 <sup>28</sup>      | 40      | Prospective | Good          | NR            | 96            | NR    | NR    | NR   | NR   |
| Dobutamine                       | Lu, et al., 2010 <sup>28</sup>      | 76      | Prospective | Good          | 87            | 82            | 77    | 90    | 4.9  | 0.16 |
| With LVH                         | Lu, et al., 2010 <sup>28</sup>      | 36      | Prospective | Good          | NR            | 69            | NR    | NR    | NR   | NR   |
| Without LVH                      | Lu, et al., 2010 <sup>28</sup>      | 40      | Prospective | Good          | NR            | 89            | NR    | NR    | NR   | NR   |
| Overall Dobutamine               | Ho, et al., 1998 <sup>67</sup>      | 51      | Prospective | Fair          | 93            | 82            | 87    | 90    | 5.12 | 0.08 |

aShin, et al. sensitivity/specificity were different in text (88/64)

Abbreviations: LR+ = positive likelihood ratio; LR- = negative likelihood ratio; N = number; NPV = negative predictive value; NR = not reported; PPV = positive predictive value



**Table 15. Summary table for SPECT**

| Predictor            | Study                               | N Women | Study Type    | Quality Score | Sensitivity % | Specificity % | PPV % | NPV % | LR+  | LR-  |
|----------------------|-------------------------------------|---------|---------------|---------------|---------------|---------------|-------|-------|------|------|
| Overall              | Ho, et al., 1998 <sup>67</sup>      | 44      | Prospective   | Fair          | 79            | 75            | 79    | 75    | 3.17 | 0.28 |
| Overall              | Vashist, et al., 2007 <sup>83</sup> | 54      | Retrospective | Fair          | 87            | 27            | 75    | 44    | 1.19 | 0.48 |
| Hispanic             | Vashist, et al., 2007 <sup>83</sup> | 16      | Retrospective | Fair          | 71.4          | 33.3          | 45.5  | 60    | 1.07 | 0.86 |
| African American     | Vashist, et al., 2007 <sup>83</sup> | 34      | Retrospective | Fair          | 90            | 20            | 86.7  | 25    | 1.12 | 0.52 |
| Overall              | Yeih, et al., 2007 <sup>50</sup>    | 51      | Prospective   | Fair          | 71            | 87            | 87    | 71    | 5.48 | 0.33 |
| Asian                | Yeih, et al., 2007 <sup>50</sup>    | 51      | Prospective   | Fair          | 71            | 87            | 87    | 71    | 5.48 | 0.33 |
| With beta blocker    | Yeih, et al., 2007 <sup>50</sup>    | 24      | Prospective   | Fair          | 67            | 78            | 83    | 58    | 3.0  | 0.43 |
| Without beta blocker | Yeih, et al., 2007 <sup>50</sup>    | 27      | Prospective   | Fair          | 77            | 93            | 91    | 81    | 10.8 | 0.25 |
| Overall              | Lu, et al., 2010 <sup>28</sup>      | 76      | Prospective   | Good          | 90            | 53            | 57    | 89    | 1.94 | 0.18 |
| With LVH             | Lu, et al., 2010 <sup>28</sup>      | 36      | Prospective   | Good          | NR            | 31            | NR    | NR    | NR   | NR   |
| Without LVH          | Lu, et al., 2010 <sup>28</sup>      | 40      | Prospective   | Good          | NR            | 66            | NR    | NR    | NR   | NR   |

Abbreviations: LR+ = positive likelihood ratio; LR- = negative likelihood ratio; LVH = left ventricular hypertrophy; CMR = cardiac magnetic resonance; N = number; NPV = negative predictive value; NR = not reported; PPV = positive predictive value

**Table 16. Summary table for CMR**

| Predictor      | Study                            | N Women | Study Type  | Quality Score | Sensitivity % | Specificity % | PPV % | NPV % | LR+  | LR-  |
|----------------|----------------------------------|---------|-------------|---------------|---------------|---------------|-------|-------|------|------|
| Overall        | Klem, et al., 2008 <sup>97</sup> | 136     | Prospective | Good          | 70            | 81            | 74    | 85    | 5.96 | 0.35 |
| ≤ 97 g LV mass | Klem, et al., 2008 <sup>97</sup> | NR      | Prospective | Good          | 69            | NR            | NR    | NR    | NR   | NR   |
| ≥ 97 g LV mass | Klem, et al., 2008 <sup>97</sup> | NR      | Prospective | Good          | 95            | NR            | NR    | NR    | NR   | NR   |

**Table 16. Summary table for CMR (continued)**

| Predictor   | Study                              | N Women              | Study Type  | Quality Score | Sensitivity %            | Specificity %          | PPV %                  | NPV %                  | LR+                        | LR-                        |
|-------------|------------------------------------|----------------------|-------------|---------------|--------------------------|------------------------|------------------------|------------------------|----------------------------|----------------------------|
| Dobutamine  | Gebker, et al., 2010 <sup>95</sup> | 183 women<br>490 men | Prospective | Good          | 85 (women)<br>86 (men)   | 86 (women)<br>83 (men) | 88 (women)<br>94 (men) | 83 (women)<br>67 (men) | 5.94 (women)<br>5.12 (men) | 0.18 (women)<br>0.17 (men) |
| With LVH    | Gebker, et al., 2010 <sup>95</sup> | 56 women<br>179 men  | Prospective | Good          | 80 (women)<br>79 (men)   | 91 (women)<br>95 (men) | 93 (women)<br>98 (men) | 73 (women)<br>56 (men) | 8.4 (women)<br>15.46 (men) | 0.22 (women)<br>0.22 (men) |
| Without LVH | Gebker, et al., 2010 <sup>95</sup> | 127 women<br>311 men | Prospective | Good          | 87.5 (women)<br>90 (men) | 84 (women)<br>78 (men) | 85 (women)<br>91 (men) | 87 (women)<br>75 (men) | 5.51 (women)<br>4.08 (men) | 0.15 (women)<br>0.12 (men) |

Abbreviations: LR+ = positive likelihood ratio; LR- = negative likelihood ratio; LVH = left ventricular hypertrophy; CMR = cardiac magnetic resonance; N = number; NPV = negative predictive value; NR = not reported; PPV = positive predictive value

## **KQ 2 Summary**

To summarize, there was insufficient available evidence to draw definitive conclusions about predictors given the small number of studies for each predictor and for each modality, as well as the combination of predictor by modality. The main predictors examined included heart size, pretest probability of CAD, race/ethnicity, postmenopausal status, and beta blocker use. No studies examined functional status, age alone, or body size. Significant variability around diagnostic accuracy existed among the studies examining each stress modality, and studies that examined heart size varied on the effect on diagnostic accuracy by stress modality.

## **Key Question 3: Use of NITs To Improve Risk Stratification, Decisionmaking, and Clinical Outcomes**

KQ 3. Is there evidence that the use of NITs (when compared with other NITs or with coronary angiography) in women improves:

KQ 3a. Risk stratification/prognostic information?

KQ 3b. Decisionmaking regarding treatment options (e.g., revascularization, optimal medical therapy)?

KQ 3c. Clinical outcomes (e.g., death, myocardial infarction, unstable angina, hospitalization, revascularization, angina relief, quality of life)?

## **Key Points**

- There were insufficient data to demonstrate that the use of specific NITs (compared with coronary angiography) routinely provided incremental risk stratification, prognostic information, or other meaningful information to improve decisionmaking.
- Compared with other NITs, stress ECG testing had higher rates of indeterminate results in women, which limited the ability to compare risk stratification with low-risk and high-risk findings from other stress imaging studies.
- There was insufficient evidence on the comparative effectiveness of different NITs to have an impact on clinical decisionmaking that leads to improved patient outcomes in women.

## **Detailed Synthesis**

For KQ 3, we examined studies that reported prognostic, outcome, or decisionmaking data comparing one NIT with another NIT or with coronary angiography in women with symptoms suspicious for CAD. We identified 13 studies,<sup>22-24,30,38,52,83,99,108-112</sup> of which 3 were considered good quality, 9 were fair quality, and 1 was poor quality. The majority of the comparative studies evaluated (n = 7) provided information on risk stratification and prognostic information. Five of these studies evaluated clinical outcomes (two studies with information on both risk stratification and clinical outcomes), and two studies were aimed at clinical decisionmaking.

In order to evaluate the ability of NITs to provide incremental information on risk stratification, prognosis, and decisionmaking, we evaluated studies that reported clinical outcomes for at least two NITs or coronary angiography. Although several studies in women

described observational cohorts of women undergoing an NIT who were followed for findings related to clinical outcomes, these studies were generally excluded since they were limited by the population risk studied, and they did not provide information on evidence for comparative effectiveness. Table 17 summarizes the findings for KQ 3a, 3b, and 3c.

### **KQ 3a: Risk Stratification and Prognostic Information**

Eight studies (two good quality, six fair)<sup>38,52,83,99,108-111</sup> provided evidence on risk stratification and prognosis. Of these studies, two<sup>108,109</sup> from the Women's Ischemia Study Evaluation (WISE) study evaluated the prognostic significance of CMR for women with suspected myocardial ischemia but without significant obstructive CAD (< 50 percent stenosis) on coronary angiography. In the study by Doyle, et al.,<sup>108</sup> two imaging variables—global magnetic resonance-myocardial perfusion imaging ratio of average peak signal-to-normalized uptake slope at stress and ejection fraction—were predictive of events including death, MI, and hospitalization for worsening angina by univariable Cox regression modeling (hazard ratio 0.516; 95% CI, 0.314 to 0.848;  $p < 0.05$  and hazard ratio of 0.949; 95% CI, 0.911 to 0.990;  $p < 0.05$ , respectively). Kaplan-Meier survival curves indicated a significant difference in time to adverse events between risk groups (log-rank 15.0,  $p < 0.0001$ ). Annualized event rates were 12 percent in the high-risk group (2 deaths and 10 hospitalizations for worsening angina) and 4 percent in the not high-risk group (2 MIs and 9 hospitalizations for worsening angina).

Johnson, et al.,<sup>109</sup> found that of the 74 women without CAD undergoing CMR, 19 percent had a cardiovascular event (1 heart failure, 12 unstable angina admission, 2 other vascular events). Women without CAD and a negative CMR had an 87-percent cumulative 3-year event-free survival rate. Women with CAD and positive (abnormal) CMR and women in the WISE study (control group, women with angiographically defined CAD) had lower event-free rates (57 percent;  $p = 0.009$ , and 52 percent;  $p < 0.0001$  respectively). Among women without CAD, the rate of hospitalization for angina was lower in those with normal CMR compared with abnormal CMR (12 percent versus 36 percent;  $p < 0.05$ ).

Two other studies<sup>52,83</sup> described the low event rate and good prognosis in patients with low-risk findings on SPECT with diagnostic validation from coronary angiography.<sup>52,83</sup> In a retrospective analysis, Vashist, et al.,<sup>83</sup> specifically studied the prognostic value of myocardial perfusion imaging in minority women (African American, Hispanic, Asian) and found that low-risk perfusion scanning signified a favorable prognosis for mortality at 2 years. Five of the 54 patients (9.3 percent) had died at 2 years (3 with intermediate-risk scan and 2 with high-risk scan). In the prospective study by Mieres, et al.,<sup>52</sup> evaluating the prognostic accuracy of SPECT in symptomatic postmenopausal women ( $n = 46$ ) with an intermediate pretest likelihood for CAD, the cumulative 3- and 5-year event-free survival was 97 and 94 percent for normal myocardial perfusion scintigraphy (MPS) compared with 60 and 48 percent for those with abnormal MPS findings ( $p < 0.0001$ ). Using ECG results for risk stratification, a negative exercise ECG was associated with 3- and 5-year event-free survival rates of 89 and 72 percent. When ECG and MPS results were included in a multivariable model, only MPS findings retained statistical significance ( $p = 0.017$ ).

The prospective analysis by Coelho-Filho, et al.,<sup>99</sup> evaluated the prognostic value of stress CMR in a consecutive group of women ( $n = 177$ ). The myocardial extent of ischemia (ISCH-SCORE) was the strongest predictor of MACE events (cardiac death and acute MI) both with univariate and multivariate analysis: hazard ratio (95% CI) 1.36 (1.23 to 1.5) and 1.49 (1.31 to 1.69) respectively. Women with evidence of ischemia also had a higher annual rate of MACE

and cardiac death compared with women without ischemia (15.1 percent and 8.2 percent versus 0.3 percent and 0 percent respectively).

Finally, three studies<sup>38,110,111</sup> demonstrated the limitations and prognostic significance of ECG compared with coronary angiography and clinical outcomes. Morise, et al.,<sup>110</sup> initially developed an exercise ECG score, including clinical and ECG variables to help risk stratify women with suspected CAD into groups of gradually increasing frequency of coronary disease and death. The score was then applied to women enrolled in the WISE study to assess the ability of the pretest and new exercise scores to stratify risk in women with low prevalence of angiographically defined CAD.<sup>111</sup> Using the pretest score, a Kaplan-Meier curve analysis of the composite endpoint (death, MI, stroke, or late revascularization) revealed a clear separation between the low-risk group and the intermediate- and high-risk groups for as long as 4 years. The intermediate- and high-risk groups were separable for as long as 1.5 years but thereafter became less clearly separable. Using the exercise population, the number of events decreased, and the Kaplan-Meier curve displayed a less clear separation and a nonsignificant difference between the curves ( $p = 0.11$ ). Using data from the WISE study, Lewis, et al.,<sup>38</sup> found that the overall sensitivity of ECG was 31 percent for obstructive CAD on coronary angiography and that the inability to perform the ECG, rather than findings on ECG, predicted MI, death, or heart failure. These data emphasize some of the limitations of using ECG in women.

### **KQ 3b: Decisionmaking for Treatment Options**

We found two studies<sup>23,112</sup> with information on clinical decisionmaking that compared different NIT strategies. The prospective analysis by Wong, et al.,<sup>112</sup> evaluated rates of referral for arteriography and revascularization according to sex. In this study, men were more likely to be referred for percutaneous transluminal coronary angioplasty or coronary artery bypass grafting than were women (59.4 percent versus 32.8 percent; odds ratio 3.0; 95% CI, 2.0 to 4.6). This difference in referral rate seemed to be linked to higher incidence of significant CAD in men (56.2 percent) compared with women (16.4 percent). When accounting for the difference in CAD incidence between men and women, there was no significant difference in revascularization rates, thus no difference in unnecessary referrals.

The study by Sanfilippo, et al.,<sup>23</sup> was an RCT of three testing strategies in women with chest pain. This fair-quality study included 158 women randomized to ECG, exercise stress ECHO, or dobutamine stress ECHO. All tests resulted in a diagnosis of cardiac chest pain or noncardiac chest pain or were indeterminate. The study found that ECG had a higher likelihood of indeterminate results, and patients with noncardiac chest pain from all modalities had low event rates. Although these two studies were informative, there remains a significant need for studies that evaluate clinical decisionmaking to improve treatment options for patients.

### **KQ 3c: Clinical Outcomes**

We found four studies that provided data on the comparative clinical outcomes for different NITs.<sup>22,24,30,38</sup> As previously stated, the study by Lewis, et al.,<sup>38</sup> identified limitations in diagnostic accuracy with ECG (overall sensitivity of 31 percent) and associated limitations in prognostic impact. Event-free survival was shorter in women who did not undergo ECG. The proportional hazards model that included ECG, CAD, and age showed estimated hazard ratios (95% CI) of 0.42 (0.18 to 0.97), 3.88 (1.72 to 8.73), and 1.01 (0.98 to 1.05) respectively ( $p = 0.0003$ ). These results indicate a decreased risk of an event for women who underwent ECG, regardless of outcome, compared with those who did not undergo ECG.

The study by Dodi, et al.,<sup>22</sup> evaluated 244 women who underwent ECG and exercise ECHO for symptoms suspicious for CAD. This study found that the prognostic information with stress ECHO (odds ratio 40) for death or MI was significantly above the effect of a positive ECG (odds ratio 3.5).

The study by Raman, et al.,<sup>30</sup> enrolled 23 women who had a positive nuclear scan (SPECT) referred for coronary angiography and who underwent dobutamine stress CMR to assess for ischemia. Followup in this study lasted  $20 \pm 8$  months, and there were no reported MI, hospitalizations, or death in their study sample.

The study by Shaw, et al.,<sup>24</sup> evaluated women with an intermediate pretest likelihood of CAD. In that study, two diagnostic strategies, exercise ECG and SPECT, had a different effect on the 2-year posttest outcomes for MACE (cardiac death, nonfatal MI) or hospital admission for acute coronary syndrome or heart failure. MACE-free survival was found to be identical (98 percent) for women randomized to the ECG or SPECT arm ( $p = 0.59$ ), and the observed 2-year MACE rate was 1.7 percent for ECG and 2.3 percent for SPECT (relative hazard, 95% CI for MACE in SPECT arm versus ECG arm was 1.3; 0.5 to 3.5;  $p = 0.59$ ). It was noted that the study was underpowered for the MACE outcome (post hoc analysis power of 15 percent at 0.05 significance level).

No articles of coronary CTA reported clinical outcomes data.

**Table 17. Summary of findings for KQ 3**

| Study                              | Study Design  | Patient Population                                  | NIT and Reference Test  | Outcomes  | Quality Score |
|------------------------------------|---|---|---|---|---------------|
| Doyle, et al., 2010 <sup>108</sup> | Single center<br>Retrospective<br>Consecutive patients<br>Enrollment: Nov 1993–Oct 1998<br>Followup: 34 ± 16 months | N overall:100<br><br>Women: 100<br><br>No known CAD | <b>NIT:</b> CMR<br><br><b>Reference test:</b><br>Coronary angiography | <b>Clinical outcomes</b><br>Adverse events: Death, MI, hospitalization for worsening angina<br>Imaging variables predictive of events by univariable Cox regression modeling were<br>Global magnetic resonance–myocardial perfusion imaging ratio of average peak signal-to-normalized uptake slope at stress, HR 0.516, 95% CI 0.314 to 0.848 (p <0.05)<br>Ejection fraction, HR 0.949, 95% CI 0.911 to 0.990 (p < 0.05)<br>Event rates<br>High-risk: 2 deaths, 10 hospitalizations for worsening angina; annualized event rate 12%<br>Low-risk: 2 MIs and 9 hospitalizations; annualized event rate 4%<br>Kaplan-Meier survival curve of time to adverse event, log-rank: 15.0, p <0.0001 | Fair          |

**Table 17. Summary of findings for KQ 3 (continued)**

| Study                                | Study Design   | Patient Population   | NIT and Reference Test  | Outcomes   | Quality Score |
|--------------------------------------|--|--|---|--|---------------|
| Johnson, et al., 2004 <sup>109</sup> | <ul style="list-style-type: none"> <li>• Multicenter</li> <li>• Prospective observational</li> <li>• Enrollment: NR</li> <li>• Followup: Median 36.5 months</li> </ul> | <p>N overall:74</p> <p>Women: 74</p> <p>Mixed population</p> | <p><b>NIT:</b> CMR</p> <p><b>Reference test:</b> Coronary angiography</p> | <p><b>Clinical outcomes</b><br/>           Death, MI, heart failure stroke, other vascular events, hospitalization for unstable angina:</p> <ul style="list-style-type: none"> <li>• In women without CAD (n = 74), 14 (19%) had cardiovascular event (0 death, 0 MI, 1 heart failure, 0 stroke, 12 unstable angina, 2 others)</li> <li>• In women without CAD and normal CMR, 87% 3-year event-free</li> <li>• In women without CAD and abnormal CMR, 57% 3-year event-free</li> </ul> <p>WISE reference women with known CAD (by coronary angiography) had 52% 3-year event-free</p> <p><b><u>Women without CAD, rate of hospitalization for angina</u></b><br/>           Normal CMR: 12%<br/>           Abnormal CMR: 36% (p &lt; 0.05)</p> <p><b><u>Repeat angiography</u></b><br/>           Normal CMR: 3%<br/>           Abnormal CMR: 21% (p &lt; 0.05)</p> | Fair          |



**Table 17. Summary of findings for KQ 3 (continued)**

| Study                               | Study Design  | Patient Population   | NIT and Reference Test   | Outcomes  | Quality Score |
|-------------------------------------|---|--|--|---|---------------|
| Mieres, et al., 2007 <sup>52</sup>  | <ul style="list-style-type: none"> <li>• Single center</li> <li>• Prospective observational</li> <li>• Consecutive patients</li> <li>• Enrollment: NR</li> <li>• Followup: 5.0 ± 3 years</li> </ul> | <p>N overall: 46</p> <p>Women: 46</p> <p>No known CAD</p>  | <p><b>NITs:</b></p> <ul style="list-style-type: none"> <li>• Exercise ECG</li> <li>• SPECT</li> </ul> <p><b>Reference test:</b><br/>Coronary angiography</p> | <p><b>Clinical outcomes</b><br/>Hospitalization for ACS, MI, or new onset or worsening angina</p> <p>Normal MPS compared with abnormal MPS</p> <p>Cumulative event-free survival at 3 and 5 years was 97% and 94% for normal MPS results compared with 60% and 48% for those with abnormal MPS findings (p &lt; 0.0001)</p> <p>Cox models looked at time to cardiovascular event as primary endpoint defined as combination of cardiovascular death, MI, increasing chest pain symptoms with definitive ECG or enzyme marker positive criteria for ACS</p> <p><b>Risk stratification/prognostic outcomes</b><br/>A negative exercise ECG was associated with 3- and 5-year event-free survival rates of 89% and 72% respectively</p> <p>Occurrence and date of the following cardiac events were noted: death, ACS, and coronary revascularization</p> <p>Coronary revascularization procedures were collected for censoring followup in the survival analysis</p> <p>Cause of death was defined as cardiovascular when occurring in the setting of a fatal MI, decompensated heart failure, or sudden cardiac death</p> <p>ACS was defined after confirmation of pain, ECG, and enzymatic criteria</p> | Fair          |
| Vashist, et al., 2007 <sup>83</sup> | <ul style="list-style-type: none"> <li>• Single center</li> <li>• Retrospective</li> <li>• Consecutive patients</li> <li>• Enrollment: NR</li> <li>• Followup: 2 years</li> </ul>                   | <p>N overall: 54</p> <p>Women: 54</p> <p>Minority women</p> <p>Mixed population—2% known CAD</p> | <p><b>NIT:</b> SPECT</p> <p><b>Reference test:</b><br/>Coronary angiography</p>  | <p><b>Clinical outcomes</b><br/>Death: N = 5 (9.3%)</p> <ul style="list-style-type: none"> <li>• 3 with intermediate-risk perfusion scan</li> <li>• 2 with high-risk perfusion scan</li> </ul> <p>Low-risk perfusion scan with low event rate over 2 years</p>  | Fair          |

**Table 17. Summary of findings for KQ 3 (continued)**

| Study                               | Study Design  | Patient Population   | NIT and Reference Test   | Outcomes  | Quality Score |
|-------------------------------------|---|--|--|---|---------------|
| Morise, et al., 2002 <sup>110</sup> | <ul style="list-style-type: none"> <li>• Multicenter</li> <li>• Prospective observational</li> <li>• Enrollment: 1981–1999</li> <li>• Followup: 2.6 years</li> </ul>  | <p>N overall: 442</p> <p>Women: 442</p> <p>No known CAD</p>  | <p><b>NIT:</b> Exercise ECG</p> <p><b>Reference test:</b> Coronary angiography</p> | <p>Study developed an exercise ECG score specifically for women.</p> <p>Factors in score for increase gradations of CAD included age (5), symptoms (2), diabetes (2), smoking (2), and estrogen status (1)</p> <p>Exercise ECG variables selected and their weights included ST depression (2), exercise heart rate (4), and Duke Angina Index (3)</p>  | Good          |
| Morise, et al., 2004 <sup>111</sup> | <ul style="list-style-type: none"> <li>• Multicenter</li> <li>• Prospective observational</li> <li>• Enrollment: NR</li> <li>• Followup: mean 3.4 years</li> </ul>  | <p>N overall: 563</p> <p>Women: 563</p> <p>No known CAD</p>  | <p><b>NIT:</b> Exercise ECG</p> <p><b>Reference test:</b> Coronary angiography</p> | <p><b>Clinical outcomes</b></p> <p>Composite death, MI, stroke, late revascularization</p> <p>Exercise scores:</p> <ul style="list-style-type: none"> <li>• Low-risk group composite endpoint in 4/83 (4.8%; annualized rate 1.4%)</li> <li>• Intermediate-risk group = 13/74 (17.6%; 5.2% annualized rate)</li> <li>• High-risk group = 4/32 (12.5%; annualized rate 3.7%, p = 0.038)</li> </ul>   | Fair          |
| Lewis, et al., 2005 <sup>38</sup>   | <ul style="list-style-type: none"> <li>• Single center</li> <li>• Prospective observational</li> <li>• Enrollment: Oct 1996–Oct 1999</li> <li>• Followup: Median 2.82 years (7 days to 5.45 years)</li> </ul> | <p>N overall: 96</p> <p>Women: 96</p> <p>Mixed population—16 known CAD (prior revascularization or MI)</p> | <p><b>NIT:</b> Exercise ECG</p> <p><b>Reference test:</b> Coronary angiography</p> | <p><b>Clinical outcomes</b></p> <p>MI: N = 5<br/>Congestive heart failure: N = 11<br/>Death: N = 12</p> <p>Overall sensitivity of exercise ECG was 31% with CAD on coronary angiography; inability to perform ECG testing, rather than findings on ECG, predicted MI, death, heart failure</p> <p>The proportional hazards model that included ECG, CAD, and age revealed estimated hazard ratios (95% CI) of 0.42 (0.18 to 0.97), 3.88 (1.72 to 8.73), and 1.01 (0.98 to 1.05), respectively (P = 0.0003)</p> <p>This indicates a decreased risk of an event for women who underwent ECG, regardless of outcome, compared with those who did not</p> | Good          |

**Table 17. Summary of findings for KQ 3 (continued)**

| Study                                  | Study Design  | Patient Population   | NIT and Reference Test  | Outcomes  | Quality Score |
|--|---|--|---|---|---------------|
| Wong, et al., 2001 <sup>112</sup>      | <ul style="list-style-type: none"> <li>• Single center</li> <li>• Retrospective observational</li> <li>• Enrollment: Dec1997–April 2000</li> <li>• Followup: NA</li> </ul>                          | <p>N overall: 1522</p> <p>Women: 601</p> <p>No known CAD</p> | <p><b>NIT:</b> Exercise ECG<br/>N = 423 (137 women 286 men)</p> <p><b>Reference test:</b><br/>Coronary angiography</p>  | <p><b>Decisionmaking for treatment</b></p> <p>Men were more likely to be referred for percutaneous transluminal coronary angioplasty or coronary artery bypass grafting than women—59.4% vs. 32.8% (OR 3.0, 95% CI 2.0 to 4.6) but men had a higher incidence of both significant and prognostic CAD (56.2% of women were found to have normal coronary arteries compared with only 16.4% of men). When this was taken into account, there was no significant difference in revascularization rates</p> | Poor          |
| Sanfilippo, et al., 2005 <sup>23</sup> | <ul style="list-style-type: none"> <li>• Multicenter</li> <li>• Prospective observational</li> <li>• Enrollment: NR</li> <li>• Followup: 28.1 ± 14.2 months</li> </ul>                              | <p>N overall: 158</p> <p>Women: 158</p> <p>No known CAD</p>  | <p><b>NIT:</b> Exercise ECG</p> <p><b>Reference test:</b><br/>Stress ECHO</p>   | <p><b>Decisionmaking for treatment</b></p> <p>Exercise ECG had higher likelihood of indeterminate results, patients with noncardiac chest pain from all modalities had low event rate</p>   | Fair          |
| Dodi, et al., 2001 <sup>22</sup>       | <ul style="list-style-type: none"> <li>• Multicenter</li> <li>• Prospective observational</li> <li>• Enrollment: Nov 1990–Oct 1996</li> <li>• Followup: 36 ± 18 months</li> </ul>                   | <p>N overall: 244</p> <p>Women: 244</p> <p>No known CAD</p>  | <p><b>NIT:</b> Exercise ECG</p> <p><b>Reference test:</b><br/>Stress ECHO</p>   | <p><b>Clinical outcomes</b></p> <p>All-cause mortality: N = 2<br/>Nonfatal MI: N = 5<br/>Unstable angina: N = 7</p> <p><b>Prognosis for death or MI</b></p> <p>Positive stress ECHO (OR 40)<br/>Positive ECG (OR 3.5)</p>   | Good          |
| Raman, et al., 2008 <sup>30</sup>      | <ul style="list-style-type: none"> <li>• Single center</li> <li>• Prospective observational</li> <li>• Consecutive patients</li> <li>• Enrollment :NR</li> <li>• Followup: 20 ± 8 months</li> </ul> | <p>N overall: 23</p> <p>Women: 23</p> <p>No known CAD</p>    | <p><b>NITs:</b></p> <ul style="list-style-type: none"> <li>• Exercise ECG</li> <li>• SPECT</li> <li>• CMR</li> </ul> <p><b>Reference test:</b><br/>Coronary angiography</p> | <p><b>Clinical outcomes</b></p> <p>MI: N = 0<br/>Hospitalization: N = 0<br/>Death: N = 0</p>  | Fair          |

**Table 17. Summary of findings for KQ 3 (continued)**

| Study                                    | Study Design   | Patient Population   | NIT and Reference Test  | Outcomes  | Quality Score |
|--|--|--|---|---|---------------|
| Coelho-Filho, et al., 2011 <sup>99</sup> | <ul style="list-style-type: none"> <li>• Single center</li> <li>• Prospective observational</li> <li>• Consecutive patients</li> <li>• Enrollment: NR</li> <li>• Followup: median 30 months</li> </ul> | <p>N overall: 405</p> <p>Women: 168</p> <p>Mixed population<br/>180 with known CAD (history of MI, PCI, or CABG)</p> | <p><b>NIT:</b> Stress CMR</p> <p><b>Reference test:</b><br/>(only in a minority of patients; N = 77)<br/>Coronary angiography</p> | <p><b><u>MACE (overall)</u></b><br/>Death: N = 15 (cardiac N = 7, noncardiac N = 8)<br/>MI: N = 7</p> <p><b><u>Ischemia (+) N = 36</u></b><br/>MACE: 15%<br/>Cardiac death: 8.2%</p> <p><b><u>Ischemia (-) N = 132</u></b><br/>MACE: 0.3%<br/>Cardiac death: 0.02%</p> <p><b><u>ISCH-SCORE</u></b><br/>MACE HR 1.49, 95% CI 1.31 to 1.69; p&lt;0.0001</p> | Fair          |

**Table 17. Summary of findings for KQ 3 (continued)**

| Study                            | Study Design   | Patient Population  | NIT and Reference Test  | Outcomes   | Quality Score |
|----------------------------------|--|---|---|--|---------------|
| Shaw, et al., 2011 <sup>24</sup> | <ul style="list-style-type: none"> <li>• Multicenter</li> <li>• RCT</li> <li>• Enrollment : NR</li> <li>• Followup: 24 months</li> </ul> | <p>N overall: 772</p> <p>Women: 772</p> <p>No known CAD</p> | <p><b>NITs:</b></p> <ul style="list-style-type: none"> <li>• Exercise ECG</li> <li>• SPECT</li> </ul> | <p><b>MACE</b></p> <p>Composite of cardiac death, nonfatal MI, or hospital admission for an acute coronary syndrome or heart failure</p> <p>MACE-free survival was identical (98%) for women randomized to the exercise ECG arm or SPECT arm (p = 0.59)</p> <p>The observed 2-year MACE rate was 1.7% for ECG and 2.3% for SPECT. The relative hazard for MACE was 1.3 (95% CI, 0.5 to 3.5) for the SPECT arm compared with the ECG arm (P = 0.59)</p> <p>Note: The study was underpowered for MACE outcome (post hoc analysis power of 15% at 0.05 significance level)</p> <p>An additional 6 women died from noncardiac causes (ECG arm, 0.5%; exercise SPECT arm, 1%; P = 0.39)</p> <p>By 6 months, 50% of enrolled women were symptom-free. By 2 years, 60% of ECG and 65% of SPECT women were symptomatic (p = 0.25)</p> <p>All Seattle Angina Questionnaire (SAQ) subscales were similar by randomized groups during followup. Cumulative incidence of worsening SAQ angina frequency or stability was 5% for both ECG and SPECT arms (p = 0.75)</p> | Fair          |

Abbreviations: CAD = coronary artery disease; CMR = cardiac magnetic resonance; ECG = exercise/stress electrocardiogram; ECHO = echocardiogram; HR = hazard ratio; MACE = major adverse cardiovascular event; MI = myocardial infarction; NIT = noninvasive technology; NA = not applicable; NR = not reported; SPECT = single proton emission computed tomography

## **KQ 3 Summary**

We identified 13 comparative studies for KQ 3 that evaluated NITs for risk stratification, prognosis, and decisionmaking affecting clinical outcomes. Two studies reported that women with abnormal CMR and normal coronary angiography had lower event-free survival rates. One study found that an abnormal SPECT resulted in a lower event-free survival rate. One study found that a negative stress ECG and diagnosis of noncardiac chest pain translated into lower event rates. Another study found that a positive stress ECHO had higher prognosis of worse cardiovascular events than a positive stress ECG. However, the studies were small and underpowered, and therefore all these findings would require significant confirmation and replication in larger studies with women. Overall, the literature identified was insufficient in demonstrating that the use of a specific NIT provided incremental risk stratification, prognostic information, or other meaningful information to improve decisionmaking and improve patient outcomes. There were specific limitations for the populations studied, including baseline risk, comparative outcomes, and relationship to diagnostic accuracy.

## **Key Question 4: Safety Concerns and Risks**

**KQ 4.** Are there significant safety concerns/risks (i.e., radiation exposure, access site complications, contrast agent-induced nephropathy, nephrogenic systemic fibrosis, anaphylaxis, arrhythmias) associated with the use of different NITs to diagnose CAD in women with symptoms suspicious for CAD?

### **Key Points**

- Data specific to women on access site complications, contrast agent-induced nephropathy, nephrogenic systemic fibrosis, or anaphylaxis associated with NITs were not reported in any of the studies included in this report.
- One study showed a significantly lower rate of supraventricular tachycardia (SVT) in women undergoing dobutamine ECHO compared with men.
- A study of adenosine thallium SPECT showed higher rates of ST depression in women compared with men.
- The results of one study suggested that lifetime attributable risk of both cancer incidence and fatal cancer incidence associated with a single coronary CTA examination was approximately twice as high in women compared with men for 50-, 60-, and 70-year old patients. These estimates are derived from projected data rather than from direct observation. The mean effective dose associated with coronary CTA in four studies ranged from 13.7 to 16.0 mSv for women and 11.1 to 16.4 for men. Radiation safety issues were not discussed in the studies that reported on other NITs.
- Other than higher mean effective radiation doses for coronary CTA studies for women compared with men, from three of the four studies reporting radiation exposure levels, there is insufficient evidence to conclude that safety concerns, risks, or radiation exposure associated with different NITs to diagnose CAD in patients with symptoms suspicious for CAD differ significantly between women and men.

## Detailed Synthesis

Tests and procedures are associated with varying degrees of risk. In the case of NITs intended to diagnose CAD in women with symptoms suspicious for CAD, physiological stress, contrast agents, and exposure to radiation may cause immediate or long-term harm. Exercise or pharmacological stressors, for example, may cause arrhythmias, acute or worsening ischemia, hypotension, or cardiac arrest. The contrast agents used in conjunction with ECHO, radionuclide myocardial perfusion imaging, CMR, and coronary CTA can cause anaphylaxis, nephrotoxicity, arrhythmias, or thromboembolism. The Food and Drug Administration (FDA) has issued a boxed warning for ECHO contrast agents that contain microscopic gas-filled spheres as well as for gadolinium-based contrast agents because of apparent risk of causing nephrogenic systemic fibrosis. CMR introduces the unique risk from metallic objects in or on the body (e.g., an aneurysm clip) causing bodily harm if moved by the magnetic forces used in CMR. Furthermore, radiation exposure associated with radionuclide myocardial perfusion imaging and coronary CTA may be carcinogenic. These safety concerns and risks may differ for women compared with men, in part because of differences in radiosensitivity between female and male reproductive organs as well as differences between reproductive organs and other organs or tissues. In the next sections, we summarize the evidence pertaining to safety concerns and risks of NITs among women, as reported and discussed in the studies included in this review (Table 18).

For KQ 4, we examined studies that reported data pertinent to safety concerns or risks associated with the use of NITs to diagnose CAD in women. We identified 13 studies,<sup>27,28,35,39,67,72,89,95,96,103,105,113,114</sup> of which 9 were good quality and 5 were fair quality.

### EKG

Five studies reported sex-specific safety data among women who underwent exercise or stress ECG and at least one other diagnostic test for CAD.<sup>27,28,35,39,67</sup> Of these studies, one was a good-quality RCT, two were good-quality prospective cohort studies, and two were fair-quality prospective cohort studies, representing a total of 413 women. Only one study<sup>28</sup> reported side effects specifically for stress ECG. In the study by Lu, et al.,<sup>28</sup> of 76 hypertensive women, rhythm disturbances were noted in 11 percent of subjects, as were frequent and severe premature ventricular contractions (PVCs). The reported adverse event rate associated with exercise ECG in this group of patients was 0 percent for symptomatic hypotension, dyspnea, nausea or vomiting, severe headache, flushing, left branch bundle block (LBBB), and supraventricular tachycardia (SVT). The other four studies reported side effects associated with ECHO but not with exercise/stress ECG.

### ECHO

Data pertaining to safety in women who underwent exercise/stress ECHO testing were reported in six studies.<sup>27,28,35,39,67,72</sup> Of these, four were good-quality prospective cohort studies, and two were fair-quality prospective cohort studies, representing a total of 513 women. One study<sup>28</sup> compared dobutamine ECHO with dipyridamole ECHO, and five studies<sup>27,35,39,67,72</sup> used dobutamine as the pharmacological stressor.

In the study by Lu, et al.,<sup>28</sup> that reported adverse events among 76 hypertensive women who underwent dobutamine ECHO, dipyridamole ECHO, and exercise ECG, the rates of adverse events associated with dobutamine ECHO were 4 percent for symptomatic hypotension, 0 percent for dyspnea, 1 percent for nausea or vomiting, 3 percent for severe headache, 0 percent for flushing, 16 percent for rhythm disturbances, 13 percent for frequent and severe PVCs, 1

percent for LBBB, and 1 percent for SVT. In contrast, the rates of adverse events associated with dipyridamole ECHO were 1 percent for symptomatic hypotension, 3 percent for dyspnea, 7 percent for nausea or vomiting, 12 percent for severe headache, 13 percent for flushing, 4 percent for rhythm disturbances, 4 percent for frequent and severe PVCs, and 0 percent each for LBBB and SVT.

In the study by Laurienzo, et al.,<sup>39</sup> that evaluated transesophageal dobutamine stress ECHO, 2 out of 84 women (2.4 percent) developed supraventricular arrhythmias, and 3 (3.6 percent) had intolerance to the probe. The study by Elhendy, et al.,<sup>71</sup> reported the symptoms and complications of dobutamine ECHO (with atropine administered as indicated) in 96 women and 210 men. Rates of reported events among the women were 5 percent for nausea, 0 percent for flushing, 2 percent for dizziness, 1 percent for anxiety, 4 percent for chills, 5 percent for headache, 1 percent for symptomatic hypotension, 38 percent for typical angina, 2 percent for SVT, 0 percent for atrial fibrillation, 1 percent for VT < 10 beats, and 0 percent for VT > 10 beats. Women experienced significantly lower rates (at the  $p < 0.05$  level) of SVT and runs of VT < 10 beats compared with men who experienced these events at rates of 9 percent and 7 percent, respectively. A study by Lewis, et al.,<sup>72</sup> that evaluated dobutamine ECHO in 92 women reported early termination of the stress test in 2 percent of patients because of VT or sustained SVT. Eight participants (9 percent) experienced dyspnea or extreme anxiety but did not require the study to be prematurely terminated, while 18 participants (20 percent) experienced mild symptoms of nausea and 8 participants (9 percent) had lightheadedness. A study of a cohort of 114 women by Lehmkuhl, et al.,<sup>27</sup> reported incidence rates of 2.6 percent for arterial hypotension, 17 percent for PVCs, and 1.7 percent for nonsustained VT with a maximum of 7 beats associated with dobutamine.

Finally, a study by Ho, et al.,<sup>67</sup> reported the following complications during dobutamine infusion for ECHO testing: frequent ventricular premature contractions (24 percent); chest pain (24 percent); palpitations (20 percent); frequent atrial premature contractions (18 percent); ST-segment change (16 percent); atrial fibrillation (2 percent); nonsustained ventricular tachycardia (2 percent); hypotension (2 percent); headache (2 percent); and yawning (2 percent).

## **SPECT**

Data pertaining to safety in women who underwent exercise/stress SPECT were reported in four prospective cohort studies<sup>28,39,67,89</sup> representing 294 women. Three studies were good-quality and one was fair-quality. The study by Lu, et al.,<sup>28</sup> evaluated technetium-99 sestamibi SPECT, the study by Ho, et al.,<sup>67</sup> compared dobutamine ECHO with SPECT, coronary angiography, and exercise ECG, and two studies, by Laurienzo, et al.,<sup>39</sup> and Mohiuddin, et al.,<sup>89</sup> evaluated thallium-201 myocardial perfusion imaging. Only one of the four studies<sup>89</sup> reported sex-specific safety data associated with SPECT. In this study of adenosine thallium-201 myocardial perfusion imaging, the rates of adverse effects of adenosine were 41 percent for flushing, 25 percent for neck or jaw pain, 30 percent for dyspnea, 12 percent for lightheadedness, 10 percent for nausea, 8 percent for headache, 4 percent for second-degree atrioventricular (AV) block, 1 percent for third-degree AV block, 48 percent for hypotension, and 20 percent for miscellaneous. Compared with men in the same study, women experienced significantly higher rates of chest pain (21 percent in men) and ST segment depression (8 percent in men) but had no significant differences in rates of other side effects.



## CMR

Two studies reported data pertaining to safety in women undergoing CMR. A study by Gebker, et al.,<sup>95</sup> reported safety data in women undergoing dobutamine stress CMR. This good-quality, prospective cohort study included 204 consecutive women and 541 men with suspected and known CAD scheduled for clinically indicated coronary angiography. In general, severe side effects likely attributable to dobutamine occurred uncommonly but tended to occur less often in women than men, with incidences of severe dyspnea of 1 percent versus 0.7 percent; severe increase in blood pressure of 0.5 percent versus 0.6 percent; paroxysmal atrial fibrillation in 1.5 percent versus 2.4 percent; incidence of ventricular tachycardia in 0.5 percent versus 0.6 percent—all in women compared with men. None of these incidences of side effects in women were statistically significantly different. The study by Merkle, et al.,<sup>96</sup> was a good-quality prospective cohort study that included 77 women who underwent both CMR and coronary angiography. This study reported no adverse events associated with adenosine infusion. Neither of the two studies reported adverse events potentially associated with CMR itself; the adverse events assessed in these two studies were limited to the pharmacological stress component of the testing procedure.

## Coronary CTA

Four studies included sex-specific data on radiation dose associated with coronary CTA.<sup>103,105,113,114</sup> All four were prospective cohort studies that compared coronary CTA with conventional coronary angiography. Of these, two were good quality and two were fair quality. Collectively, they included 486 women.

The estimated radiation exposure associated with a single 64-slice, contrast-enhanced coronary CTA was 14.4 mSv for women compared with 11.1 mSv for men in a study by Weustink, et al., (2007)<sup>113</sup> evaluating the accuracy of a 32-slice dual-source CT. A study by Dewey, et al.,<sup>103</sup> that used a 16-slice multislice CT scanner reported that the effective dose of a 16-slice coronary CTA examination was significantly higher by approximately 17 percent for women compared with men ( $13.7 \pm 1.2$  mSv versus  $11.7 \pm 0.9$  mSv,  $p < 0.001$ ). The largest contributor to dose among women were the lungs (average of 5.2 mSv, 37.8 percent of the effective dose), with breasts contributing 24.5 percent of the effective dose (3.35 mSv, on average). A study by Dharampal, et al.,<sup>105</sup> included 280 women and 636 men. Single-source CT was used for the 385 patients enrolled between July 2004 and March 2006, and dual-source CT was used for the 531 patients enrolled between April 2006 and April 2009. Unlike the previous two studies, this study found the mean effective radiation dose for single-source CT to be slightly higher in men compared with women with levels of 16.4 mSv (SD = 1.1) and 16.0 mSv (SD = 1.3) respectively ( $p = 0.002$ ). The mean effective radiation dose for dual-source CT was lower compared with single-source CT and was not significantly different between the sexes, with levels of 14.4 mSv (SD = 4.6) and 15.2 mSv (SD = 4.8) for women and men, respectively ( $p = 0.10$ ).

The fourth study, by Weustink, et al.,<sup>114</sup> included sex-specific radiation data and involved 436 symptomatic patients (301 men, 135 women; mean age of 61.6 years) who underwent both conventional coronary angiography and coronary CTA. Standard and ECG pulsing were performed in 327 and 109 patients, respectively. The authors of this study applied the Biological Effects of Ionizing Radiation (BEIR) VII approach<sup>115</sup> to estimate sex-dependent and age-dependent whole-body lifetime attributable risk of cancer incidence and mortality from a single coronary CTA examination. Risks were estimated for 50-, 60-, and 70- year-old men and women

for each of three coronary CTA techniques: no ECG pulsing, standard ECG pulsing, and optimal ECG pulsing. The findings of this study suggest that lifetime attributable risk of both cancer incidence and fatal cancer incidence was approximately double in women, compared with men for 50-, 60-, and 70-year old patients. Attributable risk was highest for no ECG pulsing and lowest for optimal ECG pulsing across all three age groups and both sexes. Attributable risk was highest for 50-year old patients and lowest for 70-year patients across all three ECG pulsing approaches and both sexes. Lifetime attributable risk of cancer associated with a single coronary CTA examination with standard ECG pulsing was estimated at approximately 0.15 percent for 60-year-old women and 0.08 percent for 60-year-old men. Lifetime attributable risk of fatal cancer associated with a single coronary CTA examination with standard ECG pulsing was estimated at approximately 0.13 percent for 60-year-old women and 0.07 percent for 60-year-old men. These estimates are derived from projected data rather than from direct observation. Of note, reproductive organs are generally more sensitive to radiation than other tissues or organs; radiation exposure to reproductive organs may therefore result in higher projected cancer risk.

**Table 18. Adverse effects of different NITs for screening of CAD in women**

| Study                          | Stress Test Modality | Comparator   | Study Type                       | N Women (Total) | Adverse Events in Women  | Adverse Events in Men | Quality |
|--------------------------------|----------------------|--|----------------------------------|-----------------|--|-----------------------|---------|
| Lu, et al., 2010 <sup>28</sup> | Exercise ECG         | Exercise MIBI scanning<br>Dobutamine ECHO<br>Dipyridamole ECHO | Prospective observational cohort | 76 (76)         | <b>Arrhythmias:</b> PVCs in 8 (11%); rhythm disturbances noted in 8 (11%). No development of LBBB or SVT<br><b>Contrast issues:</b> NA<br><b>Radiation:</b> NR<br><b>Clinical events:</b> No hypotension, dyspnea, nausea or vomiting, severe headache, flushing<br><b>Other:</b> NR | NA (women-only study) | Good    |

**Table 18. Adverse effects of different NITs for screening of CAD in women (continued)**

| Study                                | Stress Test Modality                 | Comparator                             | Study Type                       | N Women (Total) | Adverse Events in Women  | Adverse Events in Men | Quality |
|--------------------------------------|--------------------------------------|--|----------------------------------|-----------------|--|-----------------------|---------|
| Lu, et al., 2010 <sup>28</sup>       | Dobutamine ECHO<br>Dipyridamole ECHO | Exercise ECG<br>Exercise MIBI scanning | Prospective observational cohort | 76 (76)         | <p><b>Dobutamine vs. dipyridamole</b></p> <p><b>Arrhythmias:</b><br/>Rhythm disturbances: 12 (16%) vs. 3 (4%)<br/>Frequent and severe PVCs: 10 (13%) vs. 3 (4%)<br/>LBBB: 1 (1%) vs. 0<br/>SVT: 1 (1%) vs. 0</p> <p><b>Contrast issues:</b> NA<br/><b>Radiation:</b> NR<br/><b>Clinical events:</b><br/>Symptomatic hypotension: 3 (4%) vs. 1 (1%)<br/>Dyspnea: 0 vs. 2 (3%)<br/>Nausea or vomiting: 1 (1%) vs. 5 (7%)<br/>Severe headache: 2 (3%) vs. 9 (12%)<br/>Flushing: 0 vs. 10 (13%)<br/><b>Other:</b> NR</p> | NA (women-only study) | Good    |
| Lehmkuhl, et al., 2007 <sup>27</sup> | Dobutamine ECHO                      | Exercise ECG                           | Prospective observational cohort | 114 (114)       | <p><b>Arrhythmias:</b><br/>Frequent premature ventricular beats (17%)<br/>Non-sustained ventricular tachycardia of maximal 7 beats (1.7%)</p> <p><b>Contrast issues:</b> NA<br/><b>Radiation:</b> NA<br/><b>Clinical events:</b> Arterial hypotension (2.6%)<br/><b>Others:</b> NR</p>   | NA (women-only study) | Good    |

**Table 18. Adverse effects of different NITs for screening of CAD in women (continued)**

| Study                                 | Stress Test Modality            | Comparator  | Study Type                       | N Women (Total) | Adverse Events in Women  | Adverse Events in Men  | Quality |
|---------------------------------------|---------------------------------|---|----------------------------------|-----------------|--|--|---------|
| Laurienzo, et al., 1997 <sup>39</sup> | Transesophageal dobutamine ECHO | Thallium scintigraphy<br>Exercise ECG<br>Coronary angiography | Prospective observational cohort | 84 (84)         | <b>Arrhythmias:</b> SVT: 2<br><b>Contrast issues:</b> NR<br><b>Radiation:</b> NR<br><b>Clinical events:</b><br>Severe chest pain: 14<br>Severe hypertension (> 250 mm Hg): 3<br>Dobutamine-induced chest pain: 61<br><b>Other:</b><br>Extensive wall motion abnormalities: 5<br>Intolerance to probe: 3  | NA (women-only study)  | Good    |
| Elhendy, et al., 1997 <sup>35</sup>   | Dobutamine ECHO                 | Coronary angiography  | Prospective observational cohort | 96 (306)        | <b>Arrhythmias:</b><br>SVT: 2 (2)*<br>AF: 0<br>VT < 10 beats: 1 (1)*<br>VT > 10 beats: 0<br><b>*Significantly lower rates in women compared with men; others not significantly different</b><br><b>Contrast issues:</b> NR<br><b>Radiation:</b> NR<br><b>Clinical events:</b><br>Nausea: 5 (5)<br>Flushing: 0<br>Dizziness: 2 (2)<br>Anxiety: 1 (1)<br>Chills: 4 (4)<br>Headache: 5 (5)<br>Symptomatic hypotension: 1 (1)<br>Typical angina: 36 (38)<br><b>Other:</b> NR | <b>Arrhythmias:</b><br>SVT: 18 (9)*<br>AF: 1 (0.5)<br>VT < 10 beats :14 (7)*<br>VT > 10 beats: 1 (0.5)<br><b>*Significantly lower rates in women compared with men; others not significantly different</b><br><b>Contrast issues:</b> NR<br><b>Radiation:</b> NR<br><b>Clinical events:</b><br>Nausea: 4 (2)<br>Flushing: 1 (0.5)<br>Dizziness: 2 (1)<br>Anxiety: 1 (0.5)<br>Chills: 11 (5)<br>Headache: 10 (5)<br>Symptomatic hypotension: 1 (0.5)<br>Typical angina: 92 (44)<br><b>Other:</b> NR | Fair    |

**Table 18. Adverse effects of different NITs for screening of CAD in women (continued)**

| Study                             | Stress Test Modality | Comparator                                 | Study Type                       | N Women (Total) | Adverse Events in Women   | Adverse Events in Men | Quality |
|-----------------------------------|----------------------|--|----------------------------------|-----------------|---|-----------------------|---------|
| Lewis, et al., 1999 <sup>72</sup> | Dobutamine ECHO      | Coronary angiography                       | Prospective observational cohort | 92 (92)         | <b>Arrhythmias:</b><br>VT or sustained SVT requiring termination of study: 2 (2)<br>Dyspnea or extreme anxiety: 8 (9)<br><b>Contrast issues:</b> NR<br><b>Radiation:</b> NR<br><b>Clinical events:</b><br>Nausea: 18 (20)<br>Lightheadedness: 8 (9)<br>Anginal chest pain: 15 (16)<br>Hypotension: 3 (3),<br>Hypertension: 4 (4),<br>Atypical chest pain: 43 (47)<br><b>Other:</b> NR | NA (women-only study) | Good    |
| Ho, et al., 1998 <sup>67</sup>    | Dobutamine ECHO      | Exercise ECG SPECT<br>Coronary angiography | Prospective observational cohort | 51 (51)         | <b>Arrhythmias:</b><br>Frequent PVCs: 24%<br>Atrial fibrillation: 2%<br>Nonsustained VT: 2%<br>Frequent PACs: 18%<br><b>Contrast issues:</b> NR<br><b>Radiation:</b> NR<br><b>Clinical events:</b><br>Chest pain: 24%<br>Palpitations: 20%<br>Hypotension: 2%<br>Headache: 2%   | NA (women-only study) | Fair    |

**Table 18. Adverse effects of different NITs for screening of CAD in women (continued)**

| Study                                 | Stress Test Modality               | Comparator           | Study Type                       | N Women (Total) | Adverse Events in Women  | Adverse Events in Men   | Quality |
|---------------------------------------|------------------------------------|----------------------|----------------------------------|-----------------|--|---|---------|
| Mohiuddin, et al., 1996 <sup>89</sup> | Adenosine thallium 201 SPECT scans | Coronary angiography | Prospective observational cohort | 83 (202)        | <p><b>Arrhythmias:</b><br/>                     Second-degree AV block: 3 (4)<br/>                     Third-degree AV block: 1 (1.2)<br/> <b>Contrast issues:</b> NR<br/> <b>Radiation:</b> NR<br/> <b>Clinical events:</b><br/>                     Flushing: 34 (41)<br/>                     Neck or jaw pain: 21 (25)<br/>                     Dyspnea: 25 (30)<br/>                     Lightheadedness :10 (12)<br/>                     Nausea: 8 (10)<br/>                     Headache: 7 (8)<br/>                     Chest pain: 25 (30)*<br/>                     Hypotension: 40 (48)<br/> <b>Other:</b><br/>                     ST depression: 12(14)*<br/>                     Miscellaneous: 17 (20)<br/>                     *Significantly higher rates in women compared with men (p &lt; 0.05); others not significantly different</p> | <p><b>Arrhythmias:</b><br/>                     Second-degree AV block: 4 (3)<br/>                     Third-degree AV block: 1 (0.8)<br/> <b>Contrast issues:</b> NR<br/> <b>Radiation:</b> NR<br/> <b>Clinical events:</b><br/>                     Flushing: 48 (40)<br/>                     Neck or jaw pain: 33 (28)<br/>                     Dyspnea: 30 (25)<br/>                     Lightheadedness: 13 (11)<br/>                     Nausea: 12 (10)<br/>                     Headache: 9 (8)<br/>                     Chest pain: 28 (24)*<br/>                     Hypotension: 54 (45)<br/> <b>Other:</b><br/>                     ST depression 9 (8)*,<br/>                     Miscellaneous 28 (24)<br/>                     *Significantly higher rates in women compared with men (p &lt; 0.05); others not significantly different</p> | Good    |
| Gebker, et al., 2010 <sup>95</sup>    | Dobutamine stress CMR              | Coronary angiography | Prospective observational cohort | 204 (745)       | <p><b>Arrhythmias:</b><br/>                     Paroxysmal AF: 3 (1.5),<br/>                     Self-limiting VT: 1 (0.5%)<br/> <b>Contrast issues:</b> NR<br/> <b>Radiation:</b> NR<br/> <b>Clinical events:</b><br/>                     Severe chest pain 4 (2)<br/>                     Severe dyspnea 2 (1),<br/>                     Severe increase in blood pressure (&gt;240/12) 1 (0.5)<br/> <b>Other:</b> NR<br/> <b>Rates of side effects were not statistically significantly different from men in the study</b></p>  | <p><b>Arrhythmias:</b><br/>                     Paroxysmal AF: 13 (2.4)<br/>                     Self-limiting VT: 3 (0.6)<br/> <b>Contrast issues:</b> NR<br/> <b>Radiation:</b> NR<br/> <b>Clinical events:</b><br/>                     Severe chest pain 20 (3.7)<br/>                     Severe dyspnea 4 (0.7)<br/>                     Severe increase in blood pressure (&gt;240/120 mm Hg) 3 (0.6)<br/> <b>Other:</b> NR</p>  | Good    |

**Table 18. Adverse effects of different NITs for screening of CAD in women (continued)**

| Study                                 | Stress Test Modality                             | Comparator           | Study Type                       | N Women (Total) | Adverse Events in Women   | Adverse Events in Men   | Quality |
|---------------------------------------|--|----------------------|----------------------------------|-----------------|---|---|---------|
| Merkle, et al., 2010 <sup>96</sup>    | Adenosine stress CMR                             | Coronary angiography | Prospective observational cohort | 77 (256)        | <b>Arrhythmias:</b> NR<br><b>Contrast issues:</b> NR<br><b>Radiation:</b> NR<br><b>Clinical events:</b> No adverse events associated with adenosine infusion  | <b>Clinical events:</b> No adverse events associated with adenosine infusion  | Good    |
| Dewey, et al., 2008 <sup>103</sup>    | Coronary CTA (16-slice)                          | Coronary angiography | Prospective observational cohort | 50 (145)        | <b>Arrhythmias:</b> NR<br><b>Contrast issues:</b> NR<br><b>Radiation:</b> 13.7 ± 1.2 mSv*<br><b>Clinical events:</b> NR<br><b>Other:</b> NR<br><b>*Significantly higher in women compared with men (p &lt; 0.001)</b>   | <b>Arrhythmias:</b> NR<br><b>Contrast issues:</b> NR<br><b>Radiation:</b> 11.7 ± 0.9 mSv*<br><b>Clinical events:</b> NR<br><b>Other:</b> NR<br><b>*Significantly higher in women compared with men (p &lt; 0.001)</b>   | Good    |
| Weustink, et al., 2007 <sup>113</sup> | Coronary CTA (32-slice)                          | Coronary angiography | Prospective observational cohort | 21 (100)        | <b>Arrhythmias:</b> NR<br><b>Contrast issues:</b> NR<br><b>Radiation:</b> 14.4 mSv<br><b>Clinical events:</b> NR<br><b>Other:</b> NR  | <b>Arrhythmias:</b> NR<br><b>Contrast issues:</b> NR<br><b>Radiation:</b> 11.1 mSv<br><b>Clinical events:</b> NR<br><b>Other:</b> NR  | Fair    |
| Weustink, et al., 2009 <sup>114</sup> | Coronary CTA (64-slice with optimal ECG pulsing) | Coronary angiography | Prospective observational cohort | 135 (436)       | <b>Arrhythmias:</b> NR<br><b>Contrast issues:</b> NR<br><b>Radiation:</b> NR<br><b>Clinical events:</b> Approximately double lifetime attributable risk of both cancer incidence and fatal cancer incidence in women, compared with men, for 50-, 60-, and 70-year old patients<br><b>Other:</b> NR | <b>Arrhythmias:</b> NR<br><b>Contrast issues:</b> NR<br><b>Radiation:</b> NR<br><b>Clinical events:</b> Approximately double lifetime attributable risk of both cancer incidence and fatal cancer incidence in women, compared with men, for 50-, 60-, and 70-year old patients<br><b>Other:</b> NR | Good    |



**Table 18. Adverse effects of different NITs for screening of CAD in women (continued)**

| Study                                  | Stress Test Modality                            | Comparator           | Study Type                       | N Women (Total) | Adverse Events in Women  | Adverse Events in Men   | Quality |
|--|---|----------------------|----------------------------------|-----------------|--|---|---------|
| Dharampal, et al., <sup>105</sup> 2011 | Coronary CTA (single-source CT, dual-source CT) | Coronary angiography | Prospective observational cohort | 280 (916)       | <b>Arrhythmias:</b> NR<br><b>Contrast issues:</b> NR<br><b>Radiation:</b><br>Single-source CT effective radiation dose: $16.0 \pm 1.3$ mSv* (SD =1.3)<br>Dual-source CT effective radiation dose: $14.4 \pm 4.6$ mSv | <b>Arrhythmias:</b> NR<br><b>Contrast issues:</b> NR<br><b>Radiation:</b><br>Single-source CT effective radiation dose: $16.4 \pm 1.1$ mSv*<br>Dual-source CT effective radiation dose: $15.2 \pm 4.8$ mSv <sup>†</sup><br>*Significantly higher in men compared with women (p = 0.002)<br>†No significant difference in men compared with women (p = 0.10) | Fair    |

Abbreviations: AF = atrial fibrillation; CTA = computed tomography angiography; CMR = cardiac magnetic resonance imaging; ECG = electrocardiography; ECHO = echocardiography; MIBI = methoxyisobutyl; NR = not reported; NA = not applicable; PAC = premature atrial contraction; PVC = premature ventricular contraction; SVT = supraventricular tachycardia; VT = ventricular tachycardia

## **KQ 4 Summary**

Thirteen studies reported data pertinent to safety concerns or risks associated with the use of NITs to diagnose CAD in women with suspected CAD. Nine of these studies were rated good quality and four fair quality. Data pertinent to safety concerns specifically for women for a given NIT were reported in six studies for ECHO, five for coronary CTA, two for CMR, and one each for exercise/stress ECG and SPECT.

Data specific to women on access site complications, contrast agent-induced nephropathy, nephrogenic systemic fibrosis, or anaphylaxis associated with NITs were not reported in any of the studies included in this review. There was insufficient information in the extant literature to draw conclusions about sex-specific concerns about arrhythmias associated with different NITs. Total-body radiation exposure from coronary CTA examinations appeared to be higher in women compared with men; lifetime attributable risk of both cancer incidence and fatal cancer incidence associated with a single coronary CTA examination was estimated in one study to be twice as high in women compared with men. However, recent advancements in technology have reduced the radiation exposure for coronary CTA, suggesting that these estimates may not be applicable to newer testing protocols. Radiation safety issues were not discussed for NITs other than coronary CTA.

## Summary and Discussion

For this report, we conducted a systematic review of the peer-reviewed medical literature to evaluate the accuracy of different NIT modalities for diagnosing CAD in women with symptoms suspicious for CAD.

### KQ 1: Diagnostic Accuracy of NITs

For diagnostic accuracy, we identified the following number of studies for each NIT modality:

- ECG: 41 studies (13 good quality, 22 fair, 6 poor)
- ECHO: 22 studies (8 good, 13 fair, 1 poor)
- SPECT: 30 studies (10 good, 15 fair, 5 poor)
- CMR: 6 studies (5 good quality, 1 fair)
- Coronary CTA: 8 studies (4 good, 4 fair)

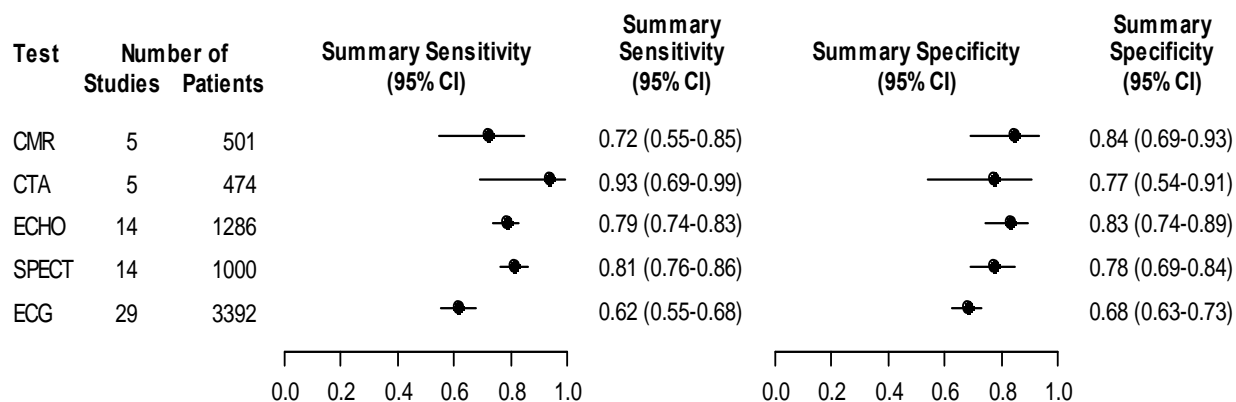
We analyzed the results by study population (no known CAD and mixed CAD populations) and by study quality (good quality rating). Table 19 and Figure 41 show the summary sensitivities and specificities for each NIT modality.

**Table 19. Summary of accuracy of NITs for diagnosing CAD in women**

| Modality     | Population       | Quality of Included Studies | Number of Studies | Number of Women | Summary Sensitivity (95% CI) | Summary Specificity (95% CI) |
|--------------|------------------|-----------------------------|-------------------|-----------------|------------------------------|------------------------------|
| ECG          | No known CAD     | All                         | 29                | 3392            | 62% (55%-68%)                | 68% (63%-73%)                |
|              |                  | Good                        | 10                | 1410            | 70% (58%-79%)                | 62% (53%-69%)                |
|              | Mixed population | All                         | 41                | 4879            | 61% (54%-67%)                | 65% (58%-72%)                |
|              |                  | Good                        | 13                | 1679            | 65% (52%-76%)                | 60% (52%-68%)                |
| ECHO         | No known CAD     | All                         | 14                | 1286            | 79% (74%-83%)                | 83% (74%-89%)                |
|              |                  | Good                        | 5                 | 561             | 79% (69%-87%)                | 85% (68%-94%)                |
|              | Mixed population | All                         | 22                | 1873            | 78% (73%-83%)                | 86% (79%-91%)                |
|              |                  | Good                        | 8                 | 807             | 77% (65%-85%)                | 89% (76%-95%)                |
| SPECT        | No known CAD     | All                         | 14                | 1000            | 81% (76%-86%)                | 78% (69%-84%)                |
|              |                  | Good                        | 4                 | 394             | 83% (52%-95%)                | 72% (37%-92%)                |
|              | Mixed population | All                         | 30                | 2146            | 82% (77%-87%)                | 81% (74%-86%)                |
|              |                  | Good                        | 10                | 982             | 82% (72%-88%)                | 79% (66%-87%)                |
| CMR          | No known CAD     | All                         | 5                 | 501             | 72% (55%-85%)                | 84% (69%-93%)                |
|              |                  | Good                        | 5                 | 501             | 72% (55%-85%)                | 84% (69%-93%)                |
|              | Mixed population | All                         | 6                 | 778             | 78% (61%-89%)                | 84% (74%-90%)                |
|              |                  | Good                        | 5                 | 610             | 76% (55%-89%)                | 84% (72%-91%)                |
| Coronary CTA | No known CAD     | All                         | 5                 | 474             | 93% (69%-99%)                | 77% (54%-91%)                |
|              |                  | Good                        | 3                 | 124             | 85% (26%-99%)                | 73% (17%-97%)                |
|              | Mixed population | All                         | 8                 | 690             | 94% (81%-98%)                | 87% (68%-96%)                |
|              |                  | Good                        | 4                 | 201             | 83% (58%-94%)                | 77% (40%-94%)                |

Abbreviations: CAD = coronary artery disease; CI = confidence interval; CMR =cardiac magnetic resonance; CTA = computed tomography angiography; ECG = exercise/stress electrocardiogram; ECHO = echocardiogram; SPECT = single proton emission computed tomography

**Figure 41. Summary of accuracy of NITs for diagnosing CAD in women with no known CAD (all studies)**



Overall, within a given modality, the summary sensitivities and specificities were similar for both types of populations (known and no known CAD) and for all studies when compared with good-quality studies. When accounting for only the good-quality studies, it appears that the diagnostic accuracy of detecting CAD in women was better (in descending order) for coronary CTA, SPECT, ECHO, CMR, and ECG. For the newer technologies (i.e., coronary CTA and CMR), more studies in women would be needed to support these findings since the 95% CIs were quite wide. In testing for a statistically significant difference between the diagnostic accuracy of testing modalities in women, our analyses determined that for women with no previously known CAD, there were differences between the performance of the available modalities ( $p < 0.001$ ). The sensitivity of ECHO and SPECT was significantly greater than that of ECG. Specificity of ECG was less than that of CMR (borderline) and of ECHO. In the subset of studies that were good-quality and where there was no known CAD in the included population, our analyses again demonstrated differences between performance of tests ( $p = 0.006$ ) with the specificity of ECG being less than that of CMR and ECHO.

To minimize the risk of spectrum bias, our primary analysis focused on women with no known CAD. We also explored mixed populations of women with known and no known CAD in sensitivity analyses. These analyses did not demonstrate a significant difference in terms of the sensitivities and specificities from our primary analysis. We also explored the prevalence of CAD across the different NIT modality studies. The mean prevalences and 95% CIs for ECG, SPECT, ECHO, CMR, and coronary CTA with the population of women with no previously known CAD was 0.41 (0.36 to 0.46), 0.44 (0.34 to 0.55), 0.43 (0.37 to 0.50), 0.26 (0.14 to 0.44), and 0.29 (0.13 to 0.54), respectively. We evaluated whether these prevalences were different across modalities using a random-effects model and did not find a statistically significant difference ( $p = 0.17$ ).

We assessed the risk of verification bias by exploring the studies in our analysis that did not complete a coronary catheterization in all of the patients who underwent the NIT. In the population of women with no previously known CAD, this represented one study of SPECT,<sup>52</sup>

one study of ECHO,<sup>79</sup> three studies of ECG,<sup>29,52,58</sup> and no studies of CMR or coronary CTA. Given the small number of total studies with this potential bias, we felt confident that our primary results were minimized for verification bias. We explored the potential for publication bias across the different modalities in our four populations of interest (studies of women with no known CAD, good-quality studies of women with no known CAD, studies of women from mixed populations, and good-quality studies of women from mixed populations). Our analyses did not provide evidence for publication bias, with our p values ranging from 0.093 to 0.95.

Table 20 shows the GRADE for the accuracy of all the NIT modalities in women with no known CAD. The number of observational studies, summary sensitivity/specificity results, and starting grade are listed for each modality. The change in the GRADE score—based on the risk of bias, consistency, directness, precision, and publication bias—is designated as “0” for no change and “-1” for a decrease due to inconsistency or imprecision. The overall strength of evidence was then determined for each modality.

**Table 20. GRADE table for accuracy of NIT modalities in women with no known CAD**

| NIT and Outcome | Quantity and Type of Evidence | Finding | Starting Grade | Decrease GRADE |             |            |           |                  | GRADE of Evidence for Outcome | Overall GRADE |
|-----------------|-------------------------------|---------|----------------|----------------|-------------|------------|-----------|------------------|-------------------------------|---------------|
|                 |                               |         |                | Risk of Bias   | Consistency | Directness | Precision | Publication Bias |                               |               |
| CMR             | 5                             |         |                |                |             |            |           |                  |                               | Low           |
| Sensitivity     |                               | 72%     | High           | 0              | -1          | 0          | -1        | 0                | Low                           |               |
| Specificity     |                               | 84%     | High           | 0              | -1          | 0          | -1        | 0                | Low                           |               |
| Coronary CTA    | 5                             |         |                |                |             |            |           |                  |                               | Low           |
| Sensitivity     |                               | 93%     | High           | 0              | -1          | 0          | -1        | 0                | Low                           |               |
| Specificity     |                               | 77%     | High           | 0              | -1          | 0          | -1        | 0                | Low                           |               |
| ECG             | 29                            |         |                |                |             |            |           |                  |                               | High          |
| Sensitivity     |                               | 62%     | High           | 0              | 0           | 0          | 0         | 0                | High                          |               |
| Specificity     |                               | 68%     | High           | 0              | 0           | 0          | 0         | 0                | High                          |               |
| ECHO            | 14                            |         |                |                |             |            |           |                  |                               | High          |
| Sensitivity     |                               | 79%     | High           | 0              | 0           | 0          | 0         | 0                | High                          |               |
| Specificity     |                               | 83%     | High           | 0              | 0           | 0          | 0         | 0                | High                          |               |
| SPECT           | 14                            |         |                |                |             |            |           |                  |                               | High          |
| Sensitivity     |                               | 81%     | High           | 0              | 0           | 0          | 0         | 0                | High                          |               |
| Specificity     |                               | 78%     | High           | 0              | 0           | 0          | 0         | 0                | High                          |               |

Abbreviations: CAD = coronary artery disease; CMR =cardiac magnetic resonance; CTA = computed tomography angiography; ECG = exercise/stress electrocardiogram; ECHO = echocardiogram; SPECT = single proton emission computed tomography

## KQ 2: Predictors of Diagnostic Accuracy

The predictors of diagnostic accuracy of the various NIT modalities in women were seldom reported. From 11 studies (4 good quality, 5 fair, and 2 poor), the predictors assessed included (1) postmenopausal women ages 55 to 64 (1 study), (2) race/ethnicity (2 studies), (3) heart size (4 studies), (4) pretest probability (3 studies), and (5) use of beta blocker medications (1 study). Despite widespread acknowledgment that patient variables can affect the accuracy of NITs, we identified no studies examining the influence of age alone, functional status, or body size on diagnostic accuracy in women. In terms of the NIT modality, we found four studies of stress ECHO, six studies of stress ECG, two studies of CMR, and four studies of SPECT that reported these predictors. There was significant variability in diagnostic accuracy as well as significant heterogeneity in the types of the predictors reported in the NIT literature. From a limited number of studies, it appeared that (1) the presence of LVH reduced the specificity of stress ECG, SPECT, and ECHO,<sup>28</sup> (2) the use of beta blocker agents reduced the specificity of stress ECG and the sensitivity and specificity of SPECT,<sup>50</sup> and (3) the positive predictive value increased as the pretest probability rose for ECG and ECHO.<sup>55</sup>

## KQ 3: Improving Risk Stratification, Decisionmaking, and Outcomes

An essential clinical question in the management of women with suspected CAD is whether NITs have the ability to provide risk stratification (low, medium, or high risk) and prognostic information (good, fair, or poor); inform decisionmaking about treatment options (medical therapy or revascularization); and affect clinical outcomes (MI, angina, quality of life, hospitalization, and death). From 13 comparative studies (3 good quality, 9 fair, 1 poor), we found 8 studies assessing risk stratification and prognostic information, 2 studies assessing decisionmaking for treatment options, and 4 studies that provided comparative clinical outcomes. Most of these findings came from subpopulations of the WISE study and assessed the prognosis and outcomes of women with normal coronary arteries on coronary angiography who underwent CMR and exercise/stress ECG. Women with normal coronary arteries by angiography and abnormal CMR or exercise ECG were more likely to have future cardiac events when compared with women with normal CMR or stress ECG findings.<sup>108,109</sup> Therefore, the study authors speculated that microvascular coronary disease could be the cause of the abnormal NIT findings and an increase in cardiovascular events. One study found that an abnormal SPECT resulted in a lower event-free survival rate. One study found that a negative stress ECG and diagnosis of noncardiac chest pain translated into lower event rates.<sup>52</sup> Another study found that a positive stress ECHO had higher prognosis of worse cardiovascular events than a positive stress ECG.<sup>22</sup> However, all these findings would require significant confirmation and replication in larger studies with women. Overall, the level of evidence was insufficient to make any conclusions about the relative utility of different NITs to provide risk stratification and prognostic information, inform decisionmaking, or impact clinical outcomes.

## KQ 4: Safety Concerns

The safety concerns and risks to women undergoing NIT procedures were underreported in the literature. We identified 13 studies (9 good quality, 4 fair) with safety data on the following modalities: (1) stress ECG (4 studies), (2) ECHO (6 studies), (3) SPECT (3 studies), (4) CMR (2 studies), and (5) coronary CTA (5 studies). One study<sup>35</sup> showed a significantly lower rate of SVT in women undergoing dobutamine ECHO. A study of adenosine thallium SPECT showed higher

rates of ST depression in women.<sup>89</sup> Four studies showed higher mean effective radiation dose or higher lifetime attributable risk of cancer incidence in women compared with men.<sup>103,105,113,114</sup> However, recent advancements in technology and revised testing protocols have reduced the radiation exposure for coronary CTA. Radiation safety issues were not discussed in other NIT modalities that employ radiation exposure (i.e., SPECT, PET, MIBI scans). The remaining studies did not show any significant adverse events in women compared with men. However, we did not find data specific to women on access site complications, contrast-agent induced nephropathy, nephrogenic systemic fibrosis, or anaphylaxis.

## Discussion

In summary, the findings of this comparative effectiveness review provide evidence for the accuracy of exercise/stress ECG, ECHO, SPECT, CMR, and coronary CTA used for diagnosing CAD in women. The diagnostic accuracy appears to be consistent over time except for the sensitivity of CMR, which appears to be increasing over time (although the large confidence intervals reflect the underlying uncertainty in this measure). We are confident that the summary statistics for ECG, ECHO, and SPECT are robust and unlikely to change with the addition of new studies based on both the number of good-quality studies comparing these modalities with coronary angiography and the tight confidence intervals. More good-quality studies comparing CMR or coronary CTA with coronary angiography in the no-known CAD population and reporting sex-based results are needed to strengthen the summary statistics for those modalities. Of note, this report focused on clinical comparative effectiveness, and so the cost of the various diagnostic strategies was not evaluated.

Decisions around performing tests (either noninvasive or invasive) in patients with symptoms suspicious for CAD revolve around first understanding the pretest probability and testing/action thresholds for patients from the AHA/ACC stable angina guidelines and appropriate use criteria for the various NIT modalities.<sup>116-119</sup> Pretest probability is classically defined by age, sex, and type of chest pain (e.g., Diamond-Forrester or CASS study), or the Duke database criteria, which adds risk factors/comorbidities.<sup>120</sup> Specifically, clinicians faced with patients who have a guideline-defined low-to-intermediate pretest probability of CAD may decide to obtain a noninvasive test, ideally with a high negative predictive value in this population and low risk of adverse events, in order to “rule out” disease. These may be patients with atypical chest pain (e.g., reflux or musculoskeletal disease) who are concerned about a heart problem and who require reassurance that their symptoms are not cardiac in origin. In contrast, in patients with high pretest probability of CAD (greater than 90 percent chance), a test with very high positive predictive value in this population and potentially more risk may be chosen since the disease of interest is thought to be present; in these cases, invasive angiography—the gold standard—is recommended by the current clinical practice guidelines. Finally, it is the spectrum of intermediate probability between 10 and 90 percent for which the clinicians must choose noninvasive tests that provide the right balance of sensitivity, specificity, and clinical risk to warrant testing. The choice of NIT may differ by clinician preference, availability, or setting (outpatient versus chest pain unit of an emergency department).

It is in this context that the findings of this report on the effectiveness of NITs in women must be considered. First, women are thought to be at lower pretest probability of CAD when evaluated in comparison with men of the same age. When comorbidities or risk factors are taken into account, the pretest likelihood increases with a higher number of comorbidities. Second, women susceptible to some of the adverse effects of testing may have poor test performance or



have higher rates of complication from invasive arterial access. Third, because of body shape and limited functional capacity, women may not obtain the same test performance that men do from noninvasive testing. Finally, because of the lack of full representation of women across the spectrum of disease, the available literature may not provide data on performance at the ends of the probability spectrum. Spectrum bias may be present since the studies we evaluated had potentially varied populations and varied disease definitions. However, this review has made a step forward in reducing the risk of spectrum bias for women by focusing on the no-known CAD subpopulation. By requiring coronary angiography as the gold standard, the pretest probability may be higher in the study population than in a routine clinical population that has a mixture of low-, intermediate-, and high-risk populations.

While readers may assume that requiring coronary angiography as the comparator would bias this report toward a higher risk CAD population, we found that the mean CAD prevalence ranged from 0.26 to 0.44; thus there was a broad spectrum of CAD prevalence in these studies. In fact, the range of CAD prevalence in this review is similar to a recent analysis of a large administrative database of patients referred for coronary angiography in which the prevalence of significant obstructive disease was 38 to 40 percent.<sup>121</sup> The patient population that does not require coronary angiography can be characterized as having symptoms with low suspicion for CAD or pretest probability of less than 10 percent (note that all included studies enrolled patients with “suspected CAD”). Thus, results from this review would not apply to patients with low pretest probability of disease (e.g., gastroesophageal reflux, musculoskeletal pain, or panic attacks) where an NIT may be performed for clinical reassurance that their symptoms are noncardiac in origin.

In general, because there are few patients with high pretest probability, most clinicians would prefer to have patients undergo one NIT prior to determining a treatment choice or referral to coronary angiography. Circumstances where patients may require more than one NIT include the detection of lesions suspicious for obstructive CAD on coronary CTA with a need to assess for ischemia from stress ECHO or SPECT prior to revascularization. Our review did not identify studies that discussed the order in which different NITs were used for evaluating CAD. In fact, *multiple testing or layered-testing strategies* are areas where significant research is needed.

The current data suggest that NITs with higher sensitivity include coronary CTA and SPECT, and stress ECHO may represent an NIT with higher specificity. Stress CMR shows emerging data that may be in the upper range for both sensitivity and specificity. Additionally, the findings also demonstrate that NIT performance in women is not as good as in men, likely due to the reasons addressed above. The accuracy may also be location or operator dependent, and thus the results of published studies conducted at highly specialized centers may not uniformly apply to those seen in routine practice. Choice of NIT—and whether to use exercise or pharmacological stress imaging—may be influenced by functional capacity, which tends to be lower in women compared with men. Of note, the accuracy data for NIT modalities in men appeared a little higher than expected, which is likely because the published literature combined the accuracy data for men and women. Taken in context, these findings support the current ACC/AHA recommendations and studies on noninvasive testing in women.

Women are more likely than men to have false positive stress tests; i.e., abnormal stress imaging with nonobstructive CAD on coronary angiography. In fact, up to 9 percent of women presenting with acute coronary syndrome will not have obstructive CAD when they undergo coronary angiography for potential PCI.<sup>122</sup> Some experts suggest that these phenomena are due

to the presence of microvascular obstruction, the incidence of which is hard to determine since there is no clear diagnostic test used to establish the diagnosis.

Currently, there is debate on whether NITs that measure heart function abnormalities (ECG abnormalities, wall motion abnormality, ischemia), including exercise ECG, stress ECHO, and cardiac nuclear imaging, are equivalent or inferior to NITs that measure anatomic abnormalities (detection of CAD) by CMR or coronary CTA. Will knowing the coronary anatomy (nonobstructive or obstructive) in symptomatic patients lead to better implementation of secondary measures—control of blood pressure, diabetes, and hyperlipidemia—to reduce future cardiac events? Or is it more important to intervene with medications and/or revascularization when ischemia is present? Though this review does not answer these important questions, we describe this evidence gap in the Future Research section.

## Limitations of This Review

Despite identifying 104 studies (110 articles) that met the inclusion criteria, this systematic review has several limitations. First, our search focused on comparator studies of the various NITs with a gold standard of coronary angiography for establishing the diagnosis of CAD in symptomatic patients. While this focus was adequate for identifying studies to assess the diagnostic accuracy of the NIT modalities in women, we found very few comparative studies that reported the influence of clinical characteristics or patient demographics on diagnostic accuracy. Few comparative studies (NIT versus coronary angiography, or NIT versus NIT) provided information on incremental risk stratification, prognostic information, or meaningful information regarding decisionmaking, and few reported the significant risks in women. Study results on these issues were reported for the total patient population and did not separate the effects by sex. Many of the included studies were single-sex (women) studies and limited our ability to fully evaluate sex differences. Also, by focusing on symptomatic patients, this report did not review the use of coronary artery calcium scoring for asymptomatic, high-risk populations.

We are aware that there are several noncomparator studies of each of the NIT modalities that address these issues in women since routine clinical care does not require two NIT modalities or an NIT modality plus coronary angiography for the diagnostic workup of suspected CAD. Given the focus on comparative effectiveness, we did not include these noncomparator studies in our review. By focusing the review on comparative studies, however, we are reducing the bias that is inherent in noncomparative studies. Noncomparative studies have selection, spectrum, and intervention biases for the following reasons: the choice of NIT is determined by the treating provider; a subset of patients with indeterminate or positive results are referred for further NIT testing or coronary angiography; and the clinical outcomes may be influenced by the medical treatments or revascularization options that are offered. Second, the sample size and low representation of women in most of the comparator studies may affect the authors' ability to analyze the results by sex, therefore reducing the number of studies reporting these findings separately. Third, most studies lacked long-term followup of the patient population, which affected our ability to find studies that reported prognostic information on how the different NITs influenced clinical outcomes. Finally, our summary of the harms and risks of NITs is limited by the lack of disclosure of periprocedural and postprocedural complications in most of the studies.

## **Conclusions**

This systematic review has provided evidence for the summary sensitivities and specificities of exercise/stress ECG, ECHO, SPECT, CMR, and coronary CTA compared with coronary angiography in women. There was limited or insufficient evidence on the influence of clinical and demographic factors on comparative diagnostic accuracy, risk stratification, prognostic information, treatment decisions, clinical outcomes, and harms from different NITs specifically in women. Modifying the search criteria to include noncomparator studies of NIT modalities may increase the number of studies that address this limitation. Table 21 summarizes the strength of supporting evidence for each KQ.

**Table 21. Summary of key findings**

| Key Question   | Strength of Evidence   | Conclusions  |
|--|--|--|
| <p><b>KQ 1: Diagnostic accuracy of NITs in women</b></p> | <p>ECG: High<br/>ECHO: High<br/>SPECT: High<br/>CMR: Low<br/>Coronary CTA: Low</p> | <p>94 studies described the diagnostic accuracy of NITs in comparison to another NIT or coronary angiography in women. Of these 94 studies, 78 studies included sufficient data to estimate the sensitivity and specificity of the NIT compared with coronary angiography.</p> <p>Summary from all studies with no known CAD:</p> <p>41 studies (13 good quality, 22 fair, 6 poor) of exercise ECG showed a summary sensitivity of 62% and specificity of 68%</p> <p>22 studies (8 good quality, 13 fair, 1 poor) of exercise/stress ECHO showed a summary sensitivity of 79% and specificity of 83%</p> <p>30 studies (10 good quality, 15 fair, 5 poor) of exercise/stress radionuclide perfusion imaging (SPECT, PET) showed a summary sensitivity of 81% and specificity of 78%</p> <p>6 studies (5 good quality, 1 fair) of CMR imaging showed a summary sensitivity of 72% and specificity of 84%</p> <p>8 studies (4 good quality, 4 fair) of coronary CTA showed a summary sensitivity of 93% and specificity 77%</p> <p>Overall, within a given modality, the summary sensitivities and specificities were similar for both types of populations (mixed populations of known and unknown CAD and no known CAD) and for all studies when compared with good-quality studies. When accounting for only the good-quality studies, it appeared that the diagnostic accuracy of detecting CAD in women with unknown CAD was better (in descending order) for coronary CTA, SPECT, ECHO, CMR, and ECG. For the newer technologies (i.e., coronary CTA and CMR), more studies in women would be needed to support these findings since the 95% CIs were quite wide.</p> <p>In testing for a statistically significant difference between the diagnostic accuracy of testing modalities in women, our analyses determined that for women with no previously known CAD, there were differences between the performance of the available modalities (<math>p &lt; 0.001</math>). The sensitivity of ECHO and SPECT was significantly higher than that of ECG. Specificity of ECG was less than that of CMR (borderline) and of ECHO. In the subset of studies that were good-quality and where there was no known CAD in the included population, our analyses again demonstrated differences between performance of tests (<math>p = 0.006</math>) with the specificity of ECG being less than that of CMR and ECHO.</p> |

**Table 21. Summary of key findings (continued)**

| Key Question  | Strength of Evidence | Conclusions   |
|---|----------------------|---|
| <b>(KQ 1 continued)</b>   |                      | Sensitivity analyses exploring mixed populations of women with known and no known CAD showed no statistically significant difference in the sensitivities and specificities from our primary analysis. An analysis exploring the prevalence of CAD across the different NIT modality studies also showed no statistically significant difference. In addition, there were very few studies (1 SPECT, 1 ECHO, and 3 ECG) that did not complete a coronary angiography in all patients who underwent the NIT; therefore the results are minimized for verification bias. Finally we found no evidence of publication bias across the different modalities in our 4 populations of interest (studies of women with no known CAD, good-quality studies of women with no known CAD, studies of women from mixed populations, and good-quality studies of women from mixed populations).  |
| <b>KQ 2: Predictors of diagnostic accuracy in women</b>                           | Insufficient         | 11 studies (4 good quality, 5 fair, 2 poor) described diagnostic accuracy, and 9 of these examined predictors of diagnostic accuracy of different NITs in women.<br>Summary:<br>The predictors assessed included (1) postmenopausal women ages 55 to 64 (1 study), (2) race/ethnicity (2 studies), (3) heart size (4 studies), (4) pretest probability (3 studies), and (5) use of beta blocker medications (1 study).<br>We identified no studies examining the influence of age alone, functional status, or body size on diagnostic accuracy in women.<br>In terms of the NIT modality, we found four studies of stress ECHO, six studies of stress ECG, two studies of CMR, and four studies of SPECT that reported these predictors.<br>Insufficient evidence was available to draw definitive conclusions about predictors given the small number of studies for each predictor and for each modality, as well as the combination of predictor by modality. |
| <b>KQ 3: Improving risk stratification, decisionmaking, and outcomes in women</b> | Insufficient         | 13 studies (3 good quality, 9 fair, 1 poor) reported prognostic, outcome, or decisionmaking data comparing one NIT with another NIT or with coronary angiography in women with symptoms suspicious for CAD.<br>Summary:<br>We found 8 studies assessing risk stratification and prognostic information, 2 studies assessing decisionmaking for treatment options, and 4 studies that provided comparative clinical outcomes.<br>There were insufficient data to demonstrate that the use of specific NITs (compared with coronary angiography) routinely provided incremental risk stratification, prognostic information, or other meaningful information to improve decisionmaking and improve patient outcomes.<br>Most findings reported in the literature would require significant confirmation and replication in larger studies with women.   |

**Table 21. Summary of key findings (continued)**

| <b>Key Question</b>          | <b>Strength of Evidence</b> | <b>Conclusions</b>   |
|------------------------------|-----------------------------|--|
| <b>KQ 4: Safety concerns</b> | Insufficient                | <p>13 studies (9 good quality, 4 fair) reported data pertinent to safety concerns or risks associated with the use of NITs to diagnose CAD in women with symptoms suspicious for CAD.</p> <p>Summary:</p> <p>Safety data were reported on the following modalities: (1) stress ECG (4 studies), (2) ECHO (6 studies), (3) SPECT (3 studies), (4) CMR (2 studies), and (5) coronary CTA (4 studies).</p> <p>Data specific to women on access site complications, contrast agent-induced nephropathy, nephrogenic systemic fibrosis, or anaphylaxis associated with NITs were not reported in any of the studies included in this report.</p> <p>Other than higher mean effective radiation doses for coronary CTA studies for women compared with men (from 3 out of 4 studies reporting radiation exposure levels), the extant literature does not provide sufficient evidence to conclude whether safety concerns, risks, or radiation exposure associated with different NITs to diagnose CAD in patients with suspected CAD differ significantly between women and men.</p> |

## Future Research

This comprehensive review of the comparative effectiveness of NIT modalities for diagnosing women with suspected CAD identified numerous gaps in evidence that would be suitable for future research and for improving the reporting of findings of NIT studies in the published literature.

**Randomized trials comparing functional versus anatomic modalities.** Almost all the studies reviewed were prospective observational studies where patients already scheduled for coronary angiography also underwent one or two NIT modalities to assess the diagnostic accuracy of the NITs. In routine clinical practice, clinicians order one type of NIT modality based on a patient's ability to exercise, test availability, and clinician preference. Exercise ECG, stress ECHO, and nuclear imaging all measure functional parameters to assess for ischemia and obstructive CAD. Newer technologies such as coronary CTA and CMR offer clinicians the ability to evaluate anatomic parameters to assess both nonobstructive and obstructive CAD. A comparison of a functional testing strategy to an anatomic testing strategy for patients with symptomatic chest pain is currently being done in two large clinical trials (PROMISE [NCT001174550] and RESCUE [NCT01262625]). The information from these clinical trials could inform how the choice of an NIT modality affects the prognosis, treatment decisions, and clinical outcomes.

**Studies assessing outcomes beyond diagnostic accuracy.** Our review found very few *comparative* NIT studies that looked at the risk stratification, prognostic information, treatment decisions, and clinical outcomes. Future studies, whether observational or controlled clinical trials, should have long-term followup of patient cohorts to assess these factors. This is important because a positive NIT result could lead to further testing to establish the diagnosis of CAD as well as lead to more attention to secondary prevention for CAD. As stated previously, multiple testing or layered-testing strategies, plus the influence on risk-factor modification (e.g., medication prescriptions and adherence), are areas where significant research is needed.

**Studies of sufficient sample size and representation of women.** Many studies assessing the comparative diagnostic accuracy of an NIT modality with another NIT modality or with coronary angiography did not present a sample size calculation for the numbers needed per group. In addition, after excluding the women-only studies, the trials with both sexes had low representation of women. In order to assess sex differences in NIT diagnostic accuracy or the impact on clinical outcomes, a sufficient sample size is required to have adequate statistical power for subgroup analyses.

**Reporting sex and CAD population subgroups separately.** From 1662 citations, we excluded 1376 (83 percent) for not reporting data on women and 615 (37 percent) for looking only at a population with known CAD. Since publication of the AHRQ report on the use of NITs in women,<sup>10,11</sup> there has been an increase in the number of studies reporting sex-based differences. We encourage more reporting of women results as well as separating the results from no known CAD and known CAD populations. One challenge we encountered in this review was that the primary data representing the numbers of TP, TN, FP, and FN were not presented in most studies and often needed to be back-calculated based on reported sensitivities and specificities and underlying disease prevalence for our quantitative synthesis. It would aid future comparisons of modalities if study authors were to report the primary data for women and men separately either within the article itself or within an online supplementary appendix.

**Assessing clinical and demographic factors that influence diagnostic accuracy.**

Clinicians are taught that clinical factors such as weight, heart size, functional status, race/ethnicity, sex, age, and menopausal status can influence the diagnostic accuracy of various NIT modalities. However, we found very few comparative studies that looked at the impact of these clinical and demographic factors on the sensitivity and specificity of NIT results. More evidence about predictors affecting diagnostic cardiac testing is needed to support or dispel these long-held notions. Additional studies of the NIT modalities to assess differing symptomatology and timing at presentation, racial differences, various risk profiles, and different settings (outpatient, inpatient, emergency room) would be help to build the evidence base needed for clinical decisionmaking.

**Reporting of risk, harms, and/or safety outcomes.** Diagnostic procedures to screen for heart disease can result in harmful clinical events (nephropathy, radiation exposure, access site complications). Systematic reporting of adverse events in publications—in total and by sex—should continue to clarify which NIT modalities are safe after they are approved for use in clinical practice.



## References

1. Mieres JH, Shaw LJ, Arai A., et al. Role of noninvasive testing in the clinical evaluation of women with suspected coronary artery disease: Consensus statement from the Cardiac Imaging Committee, Council on Clinical Cardiology, and the Cardiovascular Imaging and Intervention Committee, Council on Cardiovascular Radiology and Intervention, American Heart Association. *Circulation* 2005;111(5):682-96. PMID: 15687114
2. Lloyd-Jones D, Adams RJ, Brown TM., et al. Heart disease and stroke statistics—2010 update: a report from the American Heart Association. *Circulation* 2010;121(7):e46-e215. PMID: 20019324
3. Stangl V, Witzel V, Baumann G., et al. Current diagnostic concepts to detect coronary artery disease in women. *European Heart Journal* 2008;29(6):707-17. PMID: 18272503
4. Matchar DB, Mark DB, Patel MR., et al. Non-invasive imaging for coronary artery disease. Technology Assessment. Rockville, MD: Agency for Healthcare Research and Quality, October 2006. <http://www.cms.gov/determinationprocess/downloads/id34TA.pdf>. Accessed December 7, 2010.
5. Gibbons RJ, Balady GJ, Bricker JT., et al. ACC/AHA 2002 guideline update for exercise testing: summary article: a report of the American College of Cardiology/American Heart Association Task Force on Practice Guidelines (Committee to Update the 1997 Exercise Testing Guidelines). *Circulation* 2002;106(14):1883-92. PMID: 12356646
6. Strobeck JE, Shen JT, Singh B., et al. Comparison of a two-lead, computerized, resting ECG signal analysis device, the MultiFunction-CardioGram or MCG (a.k.a. 3DMP), to quantitative coronary angiography for the detection of relevant coronary artery stenosis (>70%)—a meta-analysis of all published trials performed and analyzed in the US. *Int J Med Sci* 2009;6(4):143-55. PMID: 19381351
7. Bluemke DA, Achenbach S, Budoff M., et al. Noninvasive coronary artery imaging: magnetic resonance angiography and multidetector computed tomography angiography: a scientific statement from the American Heart Association Committee on Cardiovascular Imaging and Intervention of the Council on Cardiovascular Radiology and Intervention, and the Councils on Clinical Cardiology and Cardiovascular Disease in the Young. *Circulation* 2008;118(5):586-606. PMID: 18586979
8. Einstein AJ. Radiation risk from coronary artery disease imaging: how do different diagnostic tests compare? *Heart* 2008;94(12):1519-21. PMID: 19011135
9. Agency for Healthcare Research and Quality. Methods Guide for Effectiveness and Comparative Effectiveness Reviews. Rockville, MD: Agency for Healthcare Research and Quality. <http://www.effectivehealthcare.ahrq.gov/index.cfm/search-for-guides-reviews-and-reports/?pageaction=displayproduct&productid=318>. Accessed August 22, 2011.
10. Grady D, Chaput L, Kristof M. Results of Systematic Review of Research on Diagnosis and Treatment of Coronary Heart Disease in Women. Evidence Report/Technology Assessment No. 80. (Prepared by the University of California, San Francisco-Stanford Evidence-based Practice Center under Contract No 290-97-0013.) AHRQ Publication No. 03-0035. Rockville,MD: Agency for Healthcare Research and Quality. May 2003. <http://archive.ahrq.gov/downloads/pub/evidence/pdf/chdwom/chdwom.pdf> Accessed July 14, 2011.

11. Grady D, Chaput L, Kristof M. Diagnosis and Treatment of Coronary Heart Disease in Women: Systematic Reviews of Evidence on Selected Topics. Evidence Report/Technology Assessment No. 81. (Prepared by the University of California, San Francisco-Stanford Evidence-based Practice Center under Contract No 290-97-0013.) AHRQ Publication No. 03-E037. Rockville, MD: Agency for Healthcare Research and Quality. May 2003.  
<http://www.ahrq.gov/downloads/pub/evidence/pdf/chdwomtop/chdwomtop.pdf>. Accessed July 14, 2011.
12. Whiting P, Rutjes AW, Reitsma JB, et al. The development of QUADAS: a tool for the quality assessment of studies of diagnostic accuracy included in systematic reviews. *BMC Med Res Methodol* 2003;3:25. PMID: 14606960
13. Atkins D, Chang SM, Gartlehner G, et al. Assessing applicability when comparing medical interventions: AHRQ and the Effective Health Care Program. *J Clin Epidemiol* 2011;64(11):1198-207. PMID: 21463926
14. Leeflang MM, Deeks JJ, Gatsonis C, et al. Systematic reviews of diagnostic test accuracy. *Ann Intern Med* 2008;149(12):889-97. PMID: 19075208
15. Blyth CR, Still HA. Binomial Confidence-Intervals. *Journal of the American Statistical Association* 1983;78(381):108-116. PMID: ISI:A1983QG01500024
16. Harbord RM, Deeks JJ, Egger M, et al. A unification of models for meta-analysis of diagnostic accuracy studies. *Biostatistics* 2007;8(2):239-51. PMID: 16698768
17. Arends LR, Hamza TH, van Houwelingen JC, et al. Bivariate random effects meta-analysis of ROC curves. *Med Decis Making* 2008;28(5):621-38. PMID: 18591542
18. Rutter CM, Gatsonis CA. Regression methods for meta-analysis of diagnostic test data. *Acad Radiol* 1995;2 Suppl 1:S48-56; discussion S65-7, S70-1 pas. PMID: 9419705
19. Deeks JJ, Macaskill P, Irwig L. The performance of tests of publication bias and other sample size effects in systematic reviews of diagnostic test accuracy was assessed. *J Clin Epidemiol* 2005;58(9):882-93. PMID: 16085191
20. Agency for Healthcare Research and Quality. Methods Guide for Medical Test Reviews [posted November 2010]. Rockville, MD. <http://effectivehealthcare.ahrq.gov/index.cfm/search-for-guides-reviews-and-reports/?pageaction=displayProduct&productID=558>. Accessed August 22, 2011.
21. Owens DK, Lohr KN, Atkins D, et al. AHRQ series paper 5: Grading the strength of a body of evidence when comparing medical interventions—Agency for Healthcare Research and Quality and the Effective Health-Care Program. *J Clin Epidemiol* 2010;63(5):513-23. PMID: 19595577
22. Dodi C, Cortigiani L, Masini M, et al. The incremental prognostic value of pharmacological stress echo over exercise electrocardiography in women with chest pain of unknown origin. *Eur Heart J* 2001;22(2):145-52. PMID: 11161916
23. Sanfilippo AJ, Abdollah H, Knott TC, et al. Stress echocardiography in the evaluation of women presenting with chest pain syndrome: a randomized, prospective comparison with electrocardiographic stress testing. *Can J Cardiol* 2005;21(5):405-12. PMID: 15861257
24. Shaw LJ, Mieres JH, Hendel RH, et al. Comparative Effectiveness of Exercise Electrocardiography With or Without Myocardial Perfusion Single Photon Emission Computed Tomography in Women With Suspected Coronary Artery Disease: Results From the What Is the Optimal Method for Ischemia Evaluation in Women (WOMEN) Trial. *Circulation* 2011. PMID: 21844080
25. Siegler JC, Rehman S, Bhumireddy GP, et al. The Accuracy of the Electrocardiogram during Exercise Stress Test Based on Heart Size. *PLoS One* 2011;6(8):e23044. PMID: 21857990
26. Doyle M, Fuisz A, Kortright E, et al. The impact of myocardial flow reserve on the detection of coronary artery disease by perfusion imaging methods: an NHLBI WISE study. *J Cardiovasc Magn Reson* 2003;5(3):475-85. PMID: 12882078
27. Lehmkuhl HB, Siniawski H, Lehmkuhl E, et al. Value and limitations of dobutamine stress echocardiography in women with suspected coronary artery disease. *Zeitschrift fur Herz-, Thorax- und Gefasschirurgie* 2007;21(6):250-258. PMID: 2008012568

28. Lu C, Lu F, Fragasso G, et al. Comparison of exercise electrocardiography, technetium-99m sestamibi single photon emission computed tomography, and dobutamine and dipyridamole echocardiography for detection of coronary artery disease in hypertensive women. *Am J Cardiol* 2010;105(9):1254-60. PMID: 20403475
29. Miller TD, Roger VL, Milavetz JJ, et al. Assessment of the exercise electrocardiogram in women versus men using tomographic myocardial perfusion imaging as the reference standard. *Am J Cardiol* 2001;87(7):868-73. PMID: 11274942
30. Raman SV, Donnally MR, McCarthy B. Dobutamine stress cardiac magnetic resonance imaging to detect myocardial ischemia in women. *Prev Cardiol* 2008;11(3):135-40. PMID: 18607148
31. Cin VG, Tartanoglu O, Duzenli A, et al. The use of basic clinical and exercise variables in postmenopausal women for the diagnosis of coronary artery disease. *Int Angiol* 2000;9(3):135-137. PMID: 2000264000
32. Hlatky MA, Pryor DB, Harrell FE, Jr, et al. Factors affecting sensitivity and specificity of exercise electrocardiography. Multivariable analysis. *Am J Med* 1984;77(1):64-71. PMID: 6741986
33. Hosokawa J, Shen JT, Imhoff M. Computerized 2-lead resting ECG analysis for the detection of relevant coronary artery stenosis in comparison with angiographic findings. *Congest Heart Fail* 2008;14(5):251-60. PMID: 18983288
34. Svart K, Lehtinen R, Nieminen T, et al. Exercise electrocardiography detection of coronary artery disease by. *Intl J Cardiol* 2010;140(2):182-188. PMID: 2010202240
35. Elhendy A, Geleijnse ML, van Domburg RT, et al. Gender differences in the accuracy of dobutamine stress echocardiography for the diagnosis of coronary artery disease. *Am J Cardiol* 1997;80(11):1414-8. PMID: 9399713
36. Yamauchi K, Simonson E, Dahl JC, et al. Sex differences in submaximal exercise tests correlation with coronary cineangiography in 133 patients. *Nagoya J Med Sci* 1985;47(1-2):67-75. PMID: 3990772
37. Friedman TD, Greene AC, Iskandrian AS, et al. Exercise thallium-201 myocardial scintigraphy in women: correlation with coronary arteriography. *Am J Cardiol* 1982;49(7):1632-7. PMID: 7081050
38. Lewis JF, McGorray S, Lin L, et al. Exercise treadmill testing using a modified exercise protocol in women with suspected myocardial ischemia: findings from the National Heart, Lung and Blood Institute-sponsored Women's Ischemia Syndrome Evaluation (WISE). *Am Heart J* 2005;149(3):527-33. PMID: 15864243
39. Laurienzo JM, Cannon RO, 3rd, Quyyumi AA, et al. Improved specificity of transesophageal dobutamine stress echocardiography compared to standard tests for evaluation of coronary artery disease in women presenting with chest pain. *Am J Cardiol* 1997;80(11):1402-7. PMID: 9399711
40. Chae SC, Heo J, Iskandrian AS, et al. Identification of extensive coronary artery disease in women by exercise single-photon emission computed tomographic (SPECT) thallium imaging. *J Am Coll Cardiol* 1993;21(6):1305-11. PMID: 8473634
41. Lewandowski M, Szwed H, Kowalik I. Searching for the optimal strategy for the diagnosis of stable coronary artery disease. Cost-effectiveness of the new algorithm. *Cardiol J* 2007;14(6):544-51. PMID: 18651520
42. Ozdemir K, Altunkeser BB, Aydin M, et al. New parameters in the interpretation of exercise testing in women: QTc dispersion and QT dispersion ratio difference. *Clin Cardiol* 2002;25(4):187-92. PMID: 12000077
43. Gentile R, Vitarelli A, Schillaci O, et al. Diagnostic accuracy and prognostic implications of stress testing for coronary artery disease in the elderly. *Ital Heart J* 2001;2(7):539-45. PMID: 11501963
44. Koide Y, Yotsukura M, Yoshino H, et al. A new coronary artery disease index of treadmill exercise electrocardiograms based on the step-up diagnostic method. *Am J Cardiol* 2001;87(2):142-7. PMID: 11152828
45. Morise AP. Are the American College of Cardiology/American Heart Association guidelines for exercise testing for suspected coronary artery disease correct? *Chest* 2000;118(2):535-41. PMID: 10936152

46. Severi S, Picano E, Michelassi C, et al. Diagnostic and prognostic value of dipyridamole echocardiography in patients with suspected coronary artery disease. Comparison with exercise electrocardiography. *Circulation* 1994;89(3):1160-73. PMID: 8124803
47. Richardson MT, Holly RG, Amsterdam EA, et al. The value of ten common exercise tolerance test measures in predicting coronary disease in symptomatic females. *Cardiology* 1995;86(3):243-8. PMID: 7614498
48. Weiner DA, Ryan TJ, McCabe CH, et al. Exercise stress testing. Correlations among history of angina, ST-segment response and prevalence of coronary-artery disease in the Coronary Artery Surgery Study (CASS). *N Engl J Med* 1979;301(5):230-5. PMID: 449990
49. Schupbach WM, Emese B, Loretan P, et al. Non-invasive diagnosis of coronary artery disease using cardiogoniometry performed at rest. *Swiss Med Wkly* 2008;138(15-16):230-8. PMID: 18431698
50. Yeih DF, Huang PJ, Ho YL. Enhanced diagnosis of coronary artery disease in women by dobutamine thallium-201 ST-segment/heart rate slope and thallium-201 myocardial SPECT. *J Formos Med Assoc* 2007;106(10):832-9. PMID: 17964962
51. Michaelides AP, Furlas CA, Chatzistamatiou EI, et al. QRS score improves diagnostic ability of treadmill exercise testing in women. *Coron Artery Dis* 2007;18(4):313-8. PMID: 17496496
52. Mieres JH, Makaryus AN, Cacciabauda JM, et al. Value of electrocardiographically gated single-photon emission computed tomographic myocardial perfusion scintigraphy in a cohort of symptomatic postmenopausal women. *Am J Cardiol* 2007;99(8):1096-9. PMID: 17437734
53. Hoiland-Carlsen PF, Johansen A, Christensen HW, et al. Usefulness of the exercise electrocardiogram in diagnosing ischemic or coronary heart disease in patients with chest pain. *Am J Cardiol* 2005;95(1):96-9. PMID: 15619400
54. Rollan MJ, San Roman JA, Vilacosta I, et al. Dobutamine stress echocardiography in the diagnosis of coronary artery disease in women with chest pain: comparison with different noninvasive tests. *Clin Cardiol* 2002;25(12):559-64. PMID: 12492125
55. Marwick TH, Anderson T, Williams MJ, et al. Exercise echocardiography is an accurate and cost-efficient technique for detection of coronary artery disease in women. *J Am Coll Cardiol* 1995;26(2):335-41. PMID: 7608432
56. Agati L, Renzi M, Sciomer S, et al. Transesophageal dipyridamole echocardiography for diagnosis of coronary artery disease. *J Am Coll Cardiol* 1992;19(4):765-70. PMID: 1545071
57. Barolsky SM, Gilbert CA, Faruqui A, et al. Differences in electrocardiographic response to exercise of women and men: a non-Bayesian factor. *Circulation* 1979;60(5):1021-7. PMID: 487535
58. Morise AP, Diamond GA. Comparison of the sensitivity and specificity of exercise electrocardiography in biased and unbiased populations of men and women. *Am Heart J* 1995;130(4):741-7. PMID: 7572581
59. Robert AR, Melin JA, Detry JM. Logistic discriminant analysis improves diagnostic accuracy of exercise testing for coronary artery disease in women. *Circulation* 1991;83(4):1202-9. PMID: 2013142
60. Guiteras P, Chaitman BR, Waters DD, et al. Diagnostic accuracy of exercise ECG lead systems in clinical subsets of women. *Circulation* 1982;65(7):1465-74. PMID: 7074802
61. Masini M, Picano E, Lattanzi F, et al. High dose dipyridamole-echocardiography test in women: correlation with exercise-electrocardiography test and coronary arteriography. *J Am Coll Cardiol* 1988;12(3):682-5. PMID: 3403825
62. Bokhari S, Shahzad A, Bergmann SR. Superiority of exercise myocardial perfusion imaging compared with the exercise ECG in the diagnosis of coronary artery disease. *Coron Artery Dis* 2008;19(6):399-404. PMID: 18955833

63. Sinha DP, Das M, Banerjee AK,, et al. Comparative study to assess whether high sensitive C-reactive protein and carotid intima media thickness improve the predictive accuracy of exercise stress testing for coronary artery disease in perimenopausal women with typical angina. *J Indian Med Assoc* 2008;106(2):86, 88, 90 passim. PMID: 18705250
64. Sketch MH, Mohiuddin SM, Lynch JD,, et al. Significant sex differences in the correlation of electrocardiographic exercise testing and coronary arteriograms. *Am J Cardiol* 1975;36(2):169-73. PMID: 1155337
65. Morise AP, Diamond GA, Detrano R,, et al. Incremental value of exercise electrocardiography and thallium-201 testing in men and women for the presence and extent of coronary artery disease. *Am Heart J* 1995;130(2):267-76. PMID: 7631606
66. Maffei E, Seitun S, Martini C,, et al. CT coronary angiography and exercise ECG in a population with chest pain and low-to-intermediate pre-test likelihood of coronary artery disease. *Heart* 2010;96(24):1973-9. PMID: 21051457
67. Ho YL, Wu CC, Huang PJ,, et al. Assessment of coronary artery disease in women by dobutamine stress echocardiography: comparison with stress thallium-201 single-photon emission computed tomography and exercise electrocardiography. *Am Heart J* 1998;135(4):655-62. PMID: 9539482
68. Santana-Boado C, Candell-Riera J, Castell-Conesa J,, et al. Diagnostic accuracy of technetium-99m-MIBI myocardial SPECT in women and men. *J Nucl Med* 1998;39(5):751-5. PMID: 9591568
69. Shin JH, Shiota T, Kim YJ,, et al. False-positive exercise echocardiograms: impact of sex and blood pressure response. *Am Heart J* 2003;146(5):914-9. PMID: 14597944
70. Bjornstad K, Aakhus S, Hatle L. Comparison of digital dipyridamole stress echocardiography and upright bicycle stress echocardiography for identification of coronary artery stenosis. *Cardiology* 1995;86(6):514-20. PMID: 7585764
71. Elhendy A, van Domburg RT, Bax JJ,, et al. Noninvasive diagnosis of coronary artery stenosis in women with limited exercise capacity: comparison of dobutamine stress echocardiography and 99mTc sestamibi single-photon emission CT. *Chest* 1998;114(4):1097-104. PMID: 9792583
72. Lewis JF, Lin L, McGorray S,, et al. Dobutamine stress echocardiography in women with chest pain. Pilot phase data from the National Heart, Lung and Blood Institute Women's Ischemia Syndrome Evaluation (WISE). *J Am Coll Cardiol* 1999;33(6):1462-8. PMID: 10334409
73. Sawada SG, Ryan T, Fineberg NS,, et al. Exercise echocardiographic detection of coronary artery disease in women. *J Am Coll Cardiol* 1989;14(6):1440-7. PMID: 2809000
74. Williams MJ, Marwick TH, O'Gorman D,, et al. Comparison of exercise echocardiography with an exercise score to diagnose coronary artery disease in women. *Am J Cardiol* 1994;74(5):435-8. PMID: 8059721
75. Dionisopoulos PN, Collins JD, Smart SC,, et al. The value of dobutamine stress echocardiography for the detection of coronary artery disease in women. *J Am Soc Echocardiogr* 1997;10(8):811-7. PMID: 9356945
76. Mazeika P, Nihoyannopoulos P, Joshi J,, et al. Uses and limitations of high dose dipyridamole stress echocardiography for evaluation of coronary artery disease. *Br Heart J* 1992;67(2):144-9. PMID: 1540434
77. Slavich GA, Guerra UP, Morocutti G,, et al. Feasibility of simultaneous Tc99m sestamibi and 2D-echo cardiac imaging during dobutamine pharmacologic stress. Preliminary results in a female population. *Int J Card Imaging* 1996;12(2):113-8. PMID: 8864790
78. Takeuchi M, Sonoda S, Miura Y,, et al. Comparative diagnostic value of dobutamine stress echocardiography and stress thallium-201 single-photon-emission computed tomography for detecting coronary artery disease in women. *Coron Artery Dis* 1996;7(11):831-5. PMID: 8993941
79. Roger VL, Pellikka PA, Bell MR,, et al. Sex and test verification bias. Impact on the diagnostic value of exercise echocardiography. *Circulation* 1997;95(2):405-10. PMID: 9008457

80. Slomka PJ, Nishina H, Abidov A,, et al. Combined quantitative supine-prone myocardial perfusion SPECT improves detection of coronary artery disease and normalcy rates in women. *J Nucl Cardiol* 2007;14(1):44-52. PMID: 17276305
81. Kaminek M, Myslivecek M, Husak V,, et al. The accuracy of myocardial perfusion SPECT imaging in the evaluation of coronary artery disease in women and men. *Nucl Med Rev Cent East Eur* 2001;4(2):69-72. PMID: 14600887
82. DePasquale EE, Nody AC, DePuey EG,, et al. Quantitative rotational thallium-201 tomography for identifying and localizing coronary artery disease. *Circulation* 1988;77(2):316-27. PMID: 3257422
83. Vashist A, Collins D, Prasad Y,, et al. Does cardiac SPECT using attenuation and scatter correction accurately predict coronary artery disease in a minority women population? *Med Sci Monit* 2007;13(9):CR386-390. PMID: 17767117
84. Gulati M, Pratap P, Kansal P,, et al. Gender differences in the value of ST-segment depression during adenosine stress testing. *Am J Cardiol* 2004;94(8):997-1002. PMID: 15476611
85. Emmett L, Iwanochko RM, Freeman MR,, et al. Reversible regional wall motion abnormalities on exercise technetium-99m-gated cardiac single photon emission computed tomography predict high-grade angiographic stenoses. *J Am Coll Cardiol* 2002;39(6):991-8. PMID: 11897441
86. Kiat H, Van Train KF, Maddahi J,, et al. Development and prospective application of quantitative 2-day stress-rest Tc-99m methoxy isobutyl isonitrile SPECT for the diagnosis of coronary artery disease. *Am Heart J* 1990;120(6 Pt 1):1255-66. PMID: 2248175
87. Taillefer R, DePuey EG, Udelson JE,, et al. Comparative diagnostic accuracy of Tl-201 and Tc-99m sestamibi SPECT imaging (perfusion and ECG-gated SPECT) in detecting coronary artery disease in women. *J Am Coll Cardiol* 1997;29(1):69-77. PMID: 8996297
88. Mak KH, Ang ES, Goh AS,, et al. Myocardial perfusion imaging with technetium-99m sestamibi SPECT in the evaluation of coronary artery disease. *Australas Radiol* 1995;39(2):112-7. PMID: 7605313
89. Mohiuddin SM, Ravage CK, Esterbrooks DJ,, et al. The comparative safety and diagnostic accuracy of adenosine myocardial perfusion imaging in women versus men. *Pharmacotherapy* 1996;16(4):646-51. PMID: 8840371
90. Hung J, Chaitman BR, Lam J,, et al. Noninvasive diagnostic test choices for the evaluation of coronary artery disease in women: a multivariate comparison of cardiac fluoroscopy, exercise electrocardiography and exercise thallium myocardial perfusion scintigraphy. *J Am Coll Cardiol* 1984;4(1):8-16. PMID: 6736458
91. Wolak A, Slomka PJ, Fish MB,, et al. Quantitative diagnostic performance of myocardial perfusion SPECT with attenuation correction in women. *J Nucl Med* 2008;49(6):915-22. PMID: 18483092
92. Elhendy A, Schinkel AF, Bax JJ,, et al. Accuracy of stress Tc-99m tetrofosmin myocardial perfusion tomography for the diagnosis and localization of coronary artery disease in women. *J Nucl Cardiol* 2006;13(5):629-34. PMID: 16945742
93. Abramson BL, Ruddy TD, deKemp RA,, et al. Stress perfusion/metabolism imaging: a pilot study for a potential new approach to the diagnosis of coronary disease in women. *J Nucl Cardiol* 2000;7(3):205-12. PMID: 10888390
94. Van Train KF, Garcia EV, Maddahi J,, et al. Multicenter trial validation for quantitative analysis of same-day rest-stress technetium-99m-sestamibi myocardial tomograms. *J Nucl Med* 1994;35(4):609-18. PMID: 8151383
95. Gebker R, Jahnke C, Hucko T,, et al. Dobutamine stress magnetic resonance imaging for the detection of coronary artery disease in women. *Heart* 2010;96(8):616-20. PMID: 19687013
96. Merkle N, Wöhrle J, Nusser T,, et al. Diagnostic performance of magnetic resonance first pass perfusion imaging is equally potent in female compared to male patients with coronary artery disease. *Clin Res Cardiol* 2010;99(1):21-8. PMID: 19756814

97. Klem I, Greulich S, Heitner JF,, et al. Value of cardiovascular magnetic resonance stress perfusion testing for the detection of coronary artery disease in women. *JACC Cardiovasc Imaging* 2008;1(4):436-45. PMID: 19356464
98. Langer C, Peterschroder A, Franzke K,, et al. Noninvasive coronary angiography focusing on calcification: multislice computed tomography compared with magnetic resonance imaging. *J Comput Assist Tomogr* 2009;33(2):179-85. PMID: 19346842
99. Coelho-Filho OR, Seabra LF, Mongeon FP,, et al. Stress Myocardial Perfusion Imaging by CMR Provides Strong Prognostic Value to Cardiac Events Regardless of Patient's Sex. *JACC Cardiovasc Imaging* 2011;4(8):850-61. PMID: 21835377
100. Dewey M, Vavere AL, Arbab-Zadeh A,, et al. Patient characteristics as predictors of image quality and diagnostic accuracy of MDCT compared with conventional coronary angiography for detecting coronary artery stenoses: CORE-64 Multicenter International Trial. *AJR Am J Roentgenol* 2010;194(1):93-102. PMID: 20028910
101. Pundziute G, Schuijff JD, Jukema JW,, et al. Gender influence on the diagnostic accuracy of 64-slice multislice computed tomography coronary angiography for detection of obstructive coronary artery disease. *Heart* 2008;94(1):48-52. PMID: 17540687
102. Jenkins SM, Johnston N, Hawkins NM,, et al. Limited clinical utility of CT coronary angiography in a district hospital setting. *QJM* 2010. PMID: 20847015
103. Dewey M, Rutsch W, Hamm B. Is there a gender difference in noninvasive coronary imaging? Multislice computed tomography for noninvasive detection of coronary stenoses. *BMC Cardiovasc Disord* 2008;8:2. PMID: 18230167
104. Shivalkar B, Goovaerts I, Salgado RA,, et al. Multislice cardiac computed tomography in symptomatic middle-aged women. *Ann Med* 2007;39(4):290-7. PMID: 17558600
105. Dharampal AS, Rossi A, Papadopoulou SL,, et al. Is there a difference in the diagnostic accuracy of computed tomography coronary angiography between women and men? *Coron Artery Dis* 2011;22(6):421-427. PMID: 2011455721
106. Burger AJ, Al-Sergani H, Wroblewski D. False positive ST segment elevation during dobutamine stress echocardiography due to left ventricular hypertrophy. *Echocardiography* 2002;19(2):103-8. PMID: 11926971
107. Gebker R, Mirelis JG, Jahnke C,, et al. Influence of left ventricular hypertrophy and geometry on diagnostic accuracy of wall motion and perfusion magnetic resonance during dobutamine stress. *Circ Cardiovasc Imaging* 2010;3(5):507-14. PMID: 20576810
108. Doyle M, Weinberg N, Pohost GM,, et al. Prognostic Value of Global MR Myocardial Perfusion Imaging in Women With Suspected Myocardial Ischemia and No Obstructive Coronary Disease Results From the NHLBI-Sponsored WISE (Women's Ischemia Syndrome Evaluation) Study. *JACC Cardiovasc Imaging* 2010;3(10):1030-6. PMID: 20947048
109. Johnson BD, Shaw LJ, Buchthal SD,, et al. Prognosis in women with myocardial ischemia in the absence of obstructive coronary disease: results from the National Institutes of Health-National Heart, Lung, and Blood Institute-Sponsored Women's Ischemia Syndrome Evaluation (WISE). *Circulation* 2004;109(24):2993-9. PMID: 15197152
110. Morise AP, Lauer MS, Froelicher VF. Development and validation of a simple exercise test score for use in women with symptoms of suspected coronary artery disease. *Am Heart J* 2002;144(5):818-25. PMID: 12422150
111. Morise AP, Olson MB, Merz CN,, et al. Validation of the accuracy of pretest and exercise test scores in women with a low prevalence of coronary disease: the NHLBI-sponsored Women's Ischemia Syndrome Evaluation (WISE) study. *Am Heart J* 2004;147(6):1085-92. PMID: 15199360
112. Wong Y, Rodwell A, Dawkins S,, et al. Sex differences in investigation results and treatment in subjects referred for investigation of chest pain. *Heart* 2001;85(2):149-52. PMID: 11156662
113. Weustink AC, Meijboom WB, Mollet NR,, et al. Reliable high-speed coronary computed tomography in symptomatic patients. *J Am Coll Cardiol* 2007;50(8):786-94. PMID: 17707184

114. Weustink AC, Mollet NR, Neeffjes LA,, et al. Preserved diagnostic performance of dual-source CT coronary angiography with reduced radiation exposure and cancer risk. *Radiology* 2009;252(1):53-60. PMID: 19451542
115. Committee to Assess Health Risks from Exposure to Low Levels of Ionizing Radiation, National Research Council. *Health Risks from Exposure to Low Levels of Ionizing Radiation: BEIR VII Phase 2*. The National Academies Press.2006. PMID:
116. Hendel RC, Patel MR, Kramer CM,, et al. ACCF/ACR/SCCT/SCMR/ASNC/NASCI/SCAI/SIR 2006 appropriateness criteria for cardiac computed tomography and cardiac magnetic resonance imaging: a report of the American College of Cardiology Foundation Quality Strategic Directions Committee Appropriateness Criteria Working Group, American College of Radiology, Society of Cardiovascular Computed Tomography, Society for Cardiovascular Magnetic Resonance, American Society of Nuclear Cardiology, North American Society for Cardiac Imaging, Society for Cardiovascular Angiography and Interventions, and Society of Interventional Radiology. *J Am Coll Cardiol* 2006;48(7):1475-97. PMID: 17010819
117. Taylor AJ, Cerqueira M, Hodgson JM,, et al. ACCF/SCCT/ACR/AHA/ASE/ASNC/NASCI/SCAI/SCMR 2010 Appropriate Use Criteria for Cardiac Computed Tomography. A Report of the American College of Cardiology Foundation Appropriate Use Criteria Task Force, the Society of Cardiovascular Computed Tomography, the American College of Radiology, the American Heart Association, the American Society of Echocardiography, the American Society of Nuclear Cardiology, the North American Society for Cardiovascular Imaging, the Society for Cardiovascular Angiography and Interventions, and the Society for Cardiovascular Magnetic Resonance. *J Cardiovasc Comput Tomogr* 2010;4(6):407 e1-33. PMID: 21232696
118. Douglas PS, Garcia MJ, Haines DE,, et al. ACCF/ASE/AHA/ASNC/HFSA/HRS/SCAI/SCCM/SCCT/SCMR 2011 Appropriate Use Criteria for Echocardiography. A Report of the American College of Cardiology Foundation Appropriate Use Criteria Task Force, American Society of Echocardiography, American Heart Association, American Society of Nuclear Cardiology, Heart Failure Society of America, Heart Rhythm Society, Society for Cardiovascular Angiography and Interventions, Society of Critical Care Medicine, Society of Cardiovascular Computed Tomography, and Society for Cardiovascular Magnetic Resonance Endorsed by the American College of Chest Physicians. *J Am Coll Cardiol* 2011;57(9):1126-66. PMID: 21349406
119. Bonow RO, Douglas PS, Buxton AE,, et al. ACCF/AHA methodology for the development of quality measures for cardiovascular technology: a report of the American College of Cardiology Foundation/American Heart Association Task Force on Performance Measures. *Circulation* 2011;124(13):1483-502. PMID: 21875906
120. Gibbons RJ, Chatterjee K, Daley J,, et al. ACC/AHA/ACP-ASIM guidelines for the management of patients with chronic stable angina: a report of the American College of Cardiology/American Heart Association Task Force on Practice Guidelines (Committee on Management of Patients With Chronic Stable Angina). *J Am Coll Cardiol* 1999;33(7):2092-197. PMID: 10362225
121. Patel MR, Peterson ED, Dai D,, et al. Low diagnostic yield of elective coronary angiography. [Erratum in: *N Engl J Med*. 2010 Jul 29;363(5):498.] *N Engl J Med* 2010;362(10):886-95. PMID: 20220183
122. Patel MR, Chen AY, Peterson ED,, et al. Prevalence, predictors, and outcomes of patients with non-ST-segment elevation myocardial infarction and insignificant coronary artery disease: results from the Can Rapid risk stratification of Unstable angina patients Suppress ADverse outcomes with Early implementation of the ACC/AHA Guidelines (CRUSADE) initiative. *Am Heart J* 2006;152(4):641-7. PMID: 16996828



## Abbreviations

|       |  |
|-------|--|
| ACC   | American College of Cardiology             |
| AHA   | American Heart Association                 |
| AV    | atrioventricular                           |
| CABG  | coronary artery bypass grafting            |
| CAD   | coronary artery disease                    |
| CMR   | cardiac magnetic resonance imaging         |
| CT    | computed tomography                        |
| CTA   | coronary computed tomography angiography   |
| ECHO  | echocardiography/echocardiogram            |
| ECG   | electrocardiography/electrocardiogram      |
| KQ    | Key Question                               |
| LBBB  | left bundle branch block                   |
| LV    | left ventricle                             |
| LVH   | left ventricular hypertrophy               |
| MI    | myocardial infarction                      |
| MIBI  | methoxyisobutyl                            |
| MPS   | myocardial perfusion scintigraphy          |
| NA    | not applicable                             |
| NIT   | noninvasive technology                     |
| NR    | not reported                               |
| PET   | positron emission tomography               |
| PVC   | premature ventricular contractions         |
| SPECT | single photon emission computed tomography |
| SVT   | supraventricular tachycardia               |
| VT    | ventricular tachycardia                    |

## Appendix A. Exact Search Strings

### PubMed® search strategy (September 12, 2011):

---

(((((diagnosis OR diagnos\* OR predict\* OR predictive value of tests OR sensitivity OR specificity) OR (sensitiv\*[Title/Abstract] OR sensitivity and specificity[MeSH Terms] OR diagnos\*[Title/Abstract] OR diagnosis[MeSH:noexp] OR diagnostic \*[MeSH:noexp] OR diagnosis, differential[MeSH:noexp] OR diagnosis[Subheading:noexp])) AND ((women OR woman OR female OR females OR sex factors) AND (((CAD[tiab]) OR (coronary artery disease[mesh] OR "coronary artery disease"[tiab] OR coronary disease[mesh] OR "coronary disease"[tiab] OR "coronary heart disease"[tiab])) OR (Chest pain OR dyspnea OR shortness of breath OR angina)) AND (((echocardiography OR echo OR cardiogram) AND ((electrocardiography OR ECG OR EKG OR electrocardio\* OR MCG OR multifunction cardiogram OR exercise test OR treadmill) OR (single photon emission computed tomography OR SPECT OR positron emission tomography OR "PET" OR myocardial perfusion imaging OR "nuclear scan" OR radionuclide imaging) OR (((cardio\* OR heart OR coronary OR cardiac) AND "Tomography, X-Ray Computed"[Mesh]) OR ("CT angiography" OR CTA OR "Cardiac Computed Tomography" OR MSCT OR Multislice computed tomography OR Multi-slice computed tomography OR MDCT OR multidetector computed tomography OR multi-detector computed tomography OR "cardiac CT" OR "Cardiovascular CT"))) OR ((cardiac OR heart OR coronary OR cardio\*) AND (magnetic resonance imaging OR MRI OR Magnetic resonance angiography OR MRA)) OR (cardiac catheterization OR angiography OR invasive coronary angiography OR heart catheterization OR coronary angiography OR "X-ray angiography" OR "Xray angiography")))) OR ((electrocardiography OR ECG OR EKG OR electrocardio\* OR MCG OR multifunction cardiogram OR exercise test OR treadmill) AND ((echocardiography OR echo OR cardiogram) OR (single photon emission computed tomography OR SPECT OR positron emission tomography OR "PET" OR myocardial perfusion imaging OR "nuclear scan" OR radionuclide imaging) OR (((cardio\* OR heart OR coronary OR cardiac) AND "Tomography, X-Ray Computed"[Mesh]) OR ("CT angiography" OR CTA OR "Cardiac Computed Tomography" OR MSCT OR Multislice computed tomography OR Multi-slice computed tomography OR MDCT OR multidetector computed tomography OR multi-detector computed tomography OR "cardiac CT" OR "Cardiovascular CT"))) OR ((cardiac OR heart OR coronary OR cardio\*) AND (magnetic resonance imaging OR MRI OR Magnetic resonance angiography OR MRA)) OR (cardiac catheterization OR angiography OR invasive coronary angiography OR heart catheterization OR coronary angiography OR "X-ray angiography" OR "Xray angiography")))) OR ((single photon emission computed tomography OR SPECT OR

positron emission tomography OR "PET" OR myocardial perfusion imaging OR "nuclear scan" OR radionuclide imaging) AND ((echocardiography OR echo OR cardiogram) OR (electrocardiography OR ECG OR EKG OR electrocardio\* OR MCG OR multifunction cardiogram OR exercise test OR treadmill) OR (((cardio\* OR heart OR coronary OR cardiac) AND "Tomography, X-Ray Computed"[Mesh]) OR ("CT angiography" OR CTA OR "Cardiac Computed Tomography" OR MSCT OR Multislice computed tomography OR Multi-slice computed tomography OR MDCT OR multidetector computed tomography OR multi-detector computed tomography OR "cardiac CT" OR "Cardiovascular CT")) OR ((cardiac OR heart OR coronary OR cardio\*) AND (magnetic resonance imaging OR MRI OR Magnetic resonance angiography OR MRA)) OR (cardiac catheterization OR angiography OR invasive coronary angiography OR heart catheterization OR coronary angiography OR "X-ray angiography" OR "Xray angiography")) OR (((cardio\* OR heart OR coronary OR cardiac) AND "Tomography, X-Ray Computed"[Mesh]) OR ("CT angiography" OR CTA OR "Cardiac Computed Tomography" OR MSCT OR Multislice computed tomography OR Multi-slice computed tomography OR MDCT OR multidetector computed tomography OR multi-detector computed tomography OR "cardiac CT" OR "Cardiovascular CT")) AND ((echocardiography OR echo OR cardiogram) OR (electrocardiography OR ECG OR EKG OR electrocardio\* OR MCG OR multifunction cardiogram OR exercise test OR treadmill) OR (single photon emission computed tomography OR SPECT OR positron emission tomography OR "PET" OR myocardial perfusion imaging OR "nuclear scan" OR radionuclide imaging) OR ((cardiac OR heart OR coronary OR cardio\*) AND (magnetic resonance imaging OR MRI OR Magnetic resonance angiography OR MRA)) OR (cardiac catheterization OR angiography OR invasive coronary angiography OR heart catheterization OR coronary angiography OR "X-ray angiography" OR "Xray angiography")) OR (((cardiac OR heart OR coronary OR cardio\*) AND (magnetic resonance imaging OR MRI OR Magnetic resonance angiography OR MRA)) AND ((echocardiography OR echo OR cardiogram) OR (electrocardiography OR ECG OR EKG OR electrocardio\* OR MCG OR multifunction cardiogram OR exercise test OR treadmill) OR (single photon emission computed tomography OR SPECT OR positron emission tomography OR "PET" OR myocardial perfusion imaging OR "nuclear scan" OR radionuclide imaging) OR (((cardio\* OR heart OR coronary OR cardiac) AND "Tomography, X-Ray Computed"[Mesh]) OR ("CT angiography" OR CTA OR "Cardiac Computed Tomography" OR MSCT OR Multislice computed tomography OR Multi-slice computed tomography OR MDCT OR multidetector computed tomography OR multi-detector computed tomography OR "cardiac CT" OR "Cardiovascular CT")) OR (cardiac catheterization OR angiography OR invasive coronary angiography OR heart catheterization OR coronary angiography OR "X-ray angiography" OR "Xray angiography")) OR ((cardiac catheterization OR angiography OR invasive coronary angiography OR heart catheterization OR coronary angiography OR "X-ray angiography" OR "Xray angiography") AND ((echocardiography OR echo OR cardiogram) OR (electrocardiography OR ECG OR EKG OR electrocardio\* OR MCG OR multifunction cardiogram OR exercise test OR treadmill) OR (single photon emission computed tomography OR SPECT OR positron emission tomography OR "PET" OR myocardial perfusion imaging OR "nuclear scan" OR radionuclide imaging) OR (((cardio\* OR heart OR coronary OR cardiac) AND "Tomography, X-Ray Computed"[Mesh]) OR ("CT angiography" OR CTA OR "Cardiac Computed Tomography" OR MSCT OR Multislice computed tomography OR Multi-slice computed tomography OR MDCT OR multidetector computed tomography OR multi-detector computed tomography OR "cardiac CT" OR "Cardiovascular CT")) OR ((cardiac OR heart OR coronary OR cardio\*) AND (magnetic

resonance imaging OR MRI OR Magnetic resonance angiography OR MRA)))))) NOT  
(Editorial[ptyp] OR Letter[ptyp] OR Case Reports[ptyp])) NOT (Animals[Mesh:noexp])

Limits:

Publication Date: 2000 – present

Language: English

## Embase® search strategy (September 12, 2011):

---

'echocardiography'/exp OR echo OR echocardiogram AND ('electrocardiography'/exp OR 'electrocardiogram'/exp OR electrocardiography OR ecg OR ekg OR electrocardio\* OR mcg OR 'multifunction cardiogram' OR ('cardiopulmonary exercise test'/exp AND 'exercise test'/exp OR exercise AND test) OR treadmill) OR ('echocardiography'/exp OR echo OR echocardiogram AND ('single photon emission computer tomography'/exp OR 'computer assisted emission tomography'/exp OR 'myocardial perfusion imaging'/exp OR 'single photon emission computed tomography' OR spect OR 'positron emission tomography' OR pet OR 'myocardial perfusion imaging' OR 'nuclear scan' OR 'radionuclide imaging' OR 'heart scintiscanning'/exp)) OR ('echocardiography'/exp OR echo OR echocardiogram AND (cardio\* OR heart OR coronary OR cardiac) AND ('computer assisted tomography'/exp OR 'computed tomographic angiography'/exp OR 'multidetector computed tomography'/exp OR 'ct angiography' OR cta OR 'cardiac computed tomography' OR 'msct' OR 'multislice computed tomography' OR 'multi-slice computed tomography' OR mdct OR 'multidetector computed tomography' OR 'multi-detector computed tomography' OR 'cardiac ct' OR 'cardiovascular ct')) OR ('echocardiography'/exp OR echo OR echocardiogram AND (cardio\* OR heart OR coronary OR cardiac) AND ('nuclear magnetic resonance imaging'/exp OR 'magnetic resonance angiography'/exp OR 'magnetic resonance imaging' OR mri OR 'magnetic resonance angiography' OR mra)) OR ('electrocardiography'/exp OR 'electrocardiogram'/exp OR electrocardiography OR ecg OR ekg OR electrocardio\* OR mcg OR 'multifunction cardiogram' OR ('cardiopulmonary exercise test'/exp AND 'exercise test'/exp OR exercise AND test) OR treadmill AND ('single photon emission computer tomography'/exp OR 'computer assisted emission tomography'/exp OR 'myocardial perfusion imaging'/exp OR 'single photon emission computed tomography' OR spect OR 'positron emission tomography' OR pet OR 'myocardial perfusion imaging' OR 'nuclear scan' OR 'radionuclide imaging' OR 'heart scintiscanning'/exp)) OR ('electrocardiography'/exp OR 'electrocardiogram'/exp OR electrocardiography OR ecg OR ekg OR electrocardio\* OR mcg OR 'multifunction cardiogram' OR ('cardiopulmonary exercise test'/exp AND 'exercise test'/exp OR exercise AND test) OR treadmill AND (cardio\* OR heart OR coronary OR cardiac) AND ('computer assisted tomography'/exp OR 'computed tomographic angiography'/exp OR 'multidetector computed tomography'/exp OR 'ct angiography' OR cta OR 'cardiac computed tomography' OR 'msct' OR 'multislice computed tomography' OR 'multi-slice computed tomography' OR mdct OR 'multidetector computed tomography' OR 'multi-detector computed tomography' OR 'cardiac ct' OR 'cardiovascular ct')) OR ('electrocardiography'/exp OR 'electrocardiogram'/exp OR electrocardiography OR ecg OR ekg OR electrocardio\* OR mcg OR 'multifunction cardiogram' OR ('cardiopulmonary exercise test'/exp AND 'exercise test'/exp OR exercise AND test) OR treadmill AND (cardio\* OR heart OR coronary OR cardiac) AND ('nuclear magnetic resonance imaging'/exp OR 'magnetic resonance angiography'/exp OR 'magnetic resonance imaging' OR mri OR 'magnetic resonance angiography' OR mra)) OR ('single photon emission computer tomography'/exp OR 'computer assisted emission tomography'/exp OR 'myocardial perfusion imaging'/exp OR 'single photon emission computed tomography' OR spect OR 'positron emission tomography' OR pet OR 'myocardial perfusion imaging' OR 'nuclear scan' OR 'radionuclide imaging' OR 'heart scintiscanning'/exp AND (cardio\* OR heart OR coronary OR cardiac) AND ('computer assisted tomography'/exp OR 'computed tomographic angiography'/exp OR 'multidetector computed tomography'/exp OR 'ct angiography' OR cta OR 'cardiac computed

tomography' OR 'msct' OR 'multislice computed tomography' OR 'multi-slice computed tomography' OR 'mdct' OR 'multidetector computed tomography' OR 'multi-detector computed tomography' OR 'cardiac ct' OR 'cardiovascular ct')) OR ('computer assisted tomography'/exp OR 'computed tomographic angiography'/exp OR 'multidetector computed tomography'/exp OR 'ct angiography' OR cta OR 'cardiac computed tomography' OR 'msct' OR 'multislice computed tomography' OR 'multi-slice computed tomography' OR 'mdct' OR 'multidetector computed tomography' OR 'multi-detector computed tomography' OR 'cardiac ct' OR 'cardiovascular ct' AND (cardio\* OR heart OR coronary OR cardiac) AND ('nuclear magnetic resonance imaging'/exp OR 'magnetic resonance angiography'/exp OR 'magnetic resonance imaging' OR mri OR 'magnetic resonance angiography' OR mra)) OR ('single photon emission computer tomography'/exp OR 'computer assisted emission tomography'/exp OR 'myocardial perfusion imaging'/exp OR 'single photon emission computed tomography' OR spect OR 'positron emission tomography' OR pet OR 'myocardial perfusion imaging' OR 'nuclear scan' OR 'radionuclide imaging' OR 'heart scintiscanning'/exp AND (cardio\* OR heart OR coronary OR cardiac) AND ('nuclear magnetic resonance imaging'/exp OR 'magnetic resonance angiography'/exp OR 'magnetic resonance imaging' OR mri OR 'magnetic resonance angiography' OR mra)) OR (cardio\* OR heart OR coronary OR cardiac AND ('nuclear magnetic resonance imaging'/exp OR 'magnetic resonance angiography'/exp OR 'magnetic resonance imaging' OR mri OR 'magnetic resonance angiography' OR mra) AND ('heart catheterization'/exp OR 'cardiac catheterization' OR angiography OR 'invasive coronary angiography' OR 'heart catheterization' OR 'coronary angiography' OR 'x-ray angiography' OR 'xray angiography')) OR ('heart catheterization'/exp OR 'cardiac catheterization' OR angiography OR 'invasive coronary angiography' OR 'heart catheterization' OR 'coronary angiography' OR 'x-ray angiography' OR 'xray angiography' AND ('echocardiography'/exp OR echo OR echocardiogram)) OR ('heart catheterization'/exp OR 'cardiac catheterization' OR angiography OR 'invasive coronary angiography' OR 'heart catheterization' OR 'coronary angiography' OR 'x-ray angiography' OR 'xray angiography' AND ('electrocardiography'/exp OR 'electrocardiogram'/exp OR electrocardiography OR ecg OR ekg OR electrocardio\* OR mcg OR 'multifunction cardiogram' OR ('cardiopulmonary exercise test'/exp AND 'exercise test'/exp OR exercise AND test) OR treadmill)) OR ('single photon emission computer tomography'/exp OR 'computer assisted emission tomography'/exp OR 'myocardial perfusion imaging'/exp OR 'single photon emission computed tomography' OR spect OR 'positron emission tomography' OR pet OR 'myocardial perfusion imaging' OR 'nuclear scan' OR 'radionuclide imaging' OR 'heart scintiscanning'/exp AND ('heart catheterization'/exp OR 'cardiac catheterization' OR angiography OR 'invasive coronary angiography' OR 'heart catheterization' OR 'coronary angiography' OR 'x-ray angiography' OR 'xray angiography')) OR (cardio\* OR heart OR coronary OR cardiac AND ('computer assisted tomography'/exp OR 'computed tomographic angiography'/exp OR 'multidetector computed tomography'/exp OR 'ct angiography' OR cta OR 'cardiac computed tomography' OR 'msct' OR 'multislice computed tomography' OR 'multi-slice computed tomography' OR 'mdct' OR 'multidetector computed tomography' OR 'multi-detector computed tomography' OR 'cardiac ct' OR 'cardiovascular ct') AND ('heart catheterization'/exp OR 'cardiac catheterization' OR angiography OR 'invasive coronary angiography' OR 'heart catheterization' OR 'coronary angiography' OR 'x-ray angiography' OR 'xray angiography')) AND ('thorax pain'/exp OR 'dyspnea'/exp OR 'angina pectoris'/exp OR 'chest pain' OR 'shortness of breath' OR angina OR dsypnea OR 'coronary artery disease'/exp OR 'ischemic heart disease'/exp OR cad OR 'coronary artery disease' OR 'coronary disease' OR 'coronary heart disease') AND ('female'/exp

OR female OR women OR woman OR females OR 'sex difference'/exp) AND (predict OR specificity OR diagnosis:lnk) AND [embase]/lim NOT [medline]/lim AND [humans]/lim AND [english]/lim AND [2000-2011]/py NOT ('case report'/exp OR 'editorial'/exp OR 'letter'/exp OR 'note'/exp)

## **Cochrane search strategy (September 12, 2011):**

-----  
[Cochrane Central Registry of Controlled Trials and Cochrane Database of Systematic Reviews]

Chest pain OR dyspnea OR shortness of breath OR angina OR CAD OR coronary artery disease OR coronary disease OR coronary heart disease

AND

((echocardiography OR echo OR cardiogram) AND ((electrocardiography OR ECG OR EKG OR electrocardio\* OR MCG OR multifunction cardiogram OR exercise test OR treadmill) OR (single photon emission computed tomography OR SPECT OR positron emission tomography OR PET OR myocardial perfusion imaging OR nuclear scan OR radionuclide imaging) OR (((cardio\* OR heart OR coronary OR cardiac) AND X-Ray computed Tomography) OR (CT angiography OR CTA OR Cardiac Computed Tomography OR MSCT OR Multislice computed tomography OR Multi-slice computed tomography OR MDCT OR multidetector computed tomography OR multi-detector computed tomography OR cardiac CT OR Cardiovascular CT)) OR ((cardiac OR heart OR coronary OR cardio\*) AND (magnetic resonance imaging OR MRI OR Magnetic resonance angiography OR MRA)) OR (cardiac catheterization OR angiography OR invasive coronary angiography OR heart catheterization OR coronary angiography OR X-ray angiography OR Xray angiography))) OR ((electrocardiography OR ECG OR EKG OR electrocardio\* OR MCG OR multifunction cardiogram OR exercise test OR treadmill) AND ((echocardiography OR echo OR cardiogram) OR (single photon emission computed tomography OR SPECT OR positron emission tomography OR PET OR myocardial perfusion imaging OR nuclear scan OR radionuclide imaging) OR (((cardio\* OR heart OR coronary OR cardiac) AND X-Ray computed Tomography) OR (CT angiography OR CTA OR Cardiac Computed Tomography OR MSCT OR Multislice computed tomography OR Multi-slice computed tomography OR MDCT OR multidetector computed tomography OR multi-detector computed tomography OR cardiac CT OR Cardiovascular CT)) OR ((cardiac OR heart OR coronary OR cardio\*) AND (magnetic resonance imaging OR MRI OR Magnetic resonance angiography OR MRA)) OR (cardiac catheterization OR angiography OR invasive coronary angiography OR heart catheterization OR coronary angiography OR X-ray angiography OR Xray angiography))) OR ((single photon emission computed tomography OR SPECT OR positron emission tomography OR PET OR myocardial perfusion imaging OR nuclear scan OR radionuclide imaging) AND ((echocardiography OR echo OR cardiogram) OR (electrocardiography OR ECG OR EKG OR electrocardio\* OR MCG OR multifunction cardiogram OR exercise test OR treadmill) OR (((cardio\* OR heart OR coronary OR cardiac) AND X-Ray computed Tomography) OR (CT angiography OR CTA OR Cardiac Computed Tomography OR MSCT OR Multislice computed tomography OR Multi-slice computed tomography OR MDCT OR multidetector computed tomography OR multi-detector computed tomography OR cardiac CT OR Cardiovascular CT)) OR ((cardiac OR heart OR coronary OR

cardio\*) AND (magnetic resonance imaging OR MRI OR Magnetic resonance angiography OR MRA)) OR (cardiac catheterization OR angiography OR invasive coronary angiography OR heart catheterization OR coronary angiography OR X-ray angiography OR Xray angiography))) OR (((cardio\* OR heart OR coronary OR cardiac) AND X-Ray computed Tomography) OR (CT angiography OR CTA OR Cardiac Computed Tomography OR MSCT OR Multislice computed tomography OR Multi-slice computed tomography OR MDCT OR multidetector computed tomography OR multi-detector computed tomography OR cardiac CT OR Cardiovascular CT)) AND ((echocardiography OR echo OR cardiogram) OR (electrocardiography OR ECG OR EKG OR electrocardio\* OR MCG OR multifunction cardiogram OR exercise test OR treadmill) OR (single photon emission computed tomography OR SPECT OR positron emission tomography OR PET OR myocardial perfusion imaging OR nuclear scan OR radionuclide imaging) OR ((cardiac OR heart OR coronary OR cardio\*) AND (magnetic resonance imaging OR MRI OR Magnetic resonance angiography OR MRA)) OR (cardiac catheterization OR angiography OR invasive coronary angiography OR heart catheterization OR coronary angiography OR X-ray angiography OR Xray angiography))) OR (((cardiac OR heart OR coronary OR cardio\*) AND (magnetic resonance imaging OR MRI OR Magnetic resonance angiography OR MRA)) AND ((echocardiography OR echo OR cardiogram) OR (electrocardiography OR ECG OR EKG OR electrocardio\* OR MCG OR multifunction cardiogram OR exercise test OR treadmill) OR (single photon emission computed tomography OR SPECT OR positron emission tomography OR PET OR myocardial perfusion imaging OR nuclear scan OR radionuclide imaging) OR (((cardio\* OR heart OR coronary OR cardiac) AND X-Ray computed Tomography) OR (CT angiography OR CTA OR Cardiac Computed Tomography OR MSCT OR Multislice computed tomography OR Multi-slice computed tomography OR MDCT OR multidetector computed tomography OR multi-detector computed tomography OR cardiac CT OR Cardiovascular CT)) OR (cardiac catheterization OR angiography OR invasive coronary angiography OR heart catheterization OR coronary angiography OR X-ray angiography OR Xray angiography))) OR ((cardiac catheterization OR angiography OR invasive coronary angiography OR heart catheterization OR coronary angiography OR X-ray angiography OR Xray angiography) AND ((echocardiography OR echo OR cardiogram) OR (electrocardiography OR ECG OR EKG OR electrocardio\* OR MCG OR multifunction cardiogram OR exercise test OR treadmill) OR (single photon emission computed tomography OR SPECT OR positron emission tomography OR PET OR myocardial perfusion imaging OR nuclear scan OR radionuclide imaging) OR (((cardio\* OR heart OR coronary OR cardiac) AND X-Ray computed Tomography) OR (CT angiography OR CTA OR Cardiac Computed Tomography OR MSCT OR Multislice computed tomography OR Multi-slice computed tomography OR MDCT OR multidetector computed tomography OR multi-detector computed tomography OR cardiac CT OR Cardiovascular CT)) OR ((cardiac OR heart OR coronary OR cardio\*) AND (magnetic resonance imaging OR MRI OR Magnetic resonance angiography OR MRA))))

AND

women OR woman OR female OR females OR sex factors

AND

diagnosis OR diagnos\* OR predict\* OR predictive value of tests OR sensitivity OR specificity OR sensitive OR diagnostic OR differential diagnosis



## Grey Literature Searches:

---

### **ClinicalTrials.gov**

searched: 12-6-2010

noninvasive [ALL-FIELDS] AND coronary artery disease [DISEASE] AND ( NOT "Male" )  
[GENDER] AND "completed" [SUMMARY-STATUS]

### **metaRegister of Controlled Trials (mRCT)**

searched: 12-6-2010

(noninvasive OR non-invasive OR "non invasive") AND ("coronary artery disease" OR CAD)  
[no results]

coronary artery disease [26 results scanned for completed trials related to diagnosis - no results]

### **ClinicalStudyResults.org**

searched: 12-6-2010

CAD OR "coronary artery disease" [no results]

### **WHO: International Clinical Trials Registry Platform Search Portal**

searched: 12-6-2010

(noninvasive OR non-invasive OR "non invasive") in Title  
AND

("coronary artery disease" OR CAD) in Indication  
scanned for completed trials related to diagnosis

### **CSA Conference Papers Index**

searched: 12-6-2010

Search Query #2 KW=(noninvasive or non-invasive or (non invasive)) and KW=(cad or  
(coronary artery disease)) and KW=(diagnosis or detection or screening) (Copy Query)

24 Published Works results found in Conference Papers Index

Date Range: Earliest to 2011

### **Scopus**

searched: 12-6-2010

Your query: (TITLE-ABS-KEY(screening OR detection OR diagnosis OR assessment)) AND  
((TITLE-ABS-KEY(noninvasive OR "non invasive" OR non-invasive)) AND (TITLE-ABS-  
KEY("coronary artery disease" OR cad))) AND (LIMIT-TO(DOCTYPE, "cp"))

# Appendix B. Data Abstraction Elements

## I. Study Characteristics

- Study dates
- Study sites
- Geographic location
- Funding source
- Study design
- If discernable: Is this article known to be a report of data from a population discussed in another article?
  - If Yes, note the primary publication for the study by entering the citation information or study identifier (trial name, acronym, or NCT number). Citation information is preferred.
- Testing setting (select all that apply)
  - Emergency Department/ Chest Pain Unit
  - Outpatient
  - Inpatient
  - Other (specify)
  - Not Reported/Unclear
- Duration of longest follow-up after completion of final test. Enter with units (days, weeks, months). NR if Not Reported. Enter NA if Not Applicable (i.e. if the study did not include a follow-up period).
- Was screening and enrollment consecutive?
- Inclusion criteria: Copy/paste inclusion criteria as reported in the article.
- Exclusion criteria: Copy/paste exclusion criteria as reported in the article.
- Study Enrollment
  - Total population
    - Number of subjects enrolled
    - Number of subjects with known CAD
    - Number of subjects without known CAD
    - If applicable, enter the definition of known CAD
  - Female
    - Number of subjects enrolled
    - Number of subjects with known CAD
    - Number of subjects without known CAD
    - If applicable, enter the definition of known CAD
  - Male
    - Number of subjects enrolled
    - Number of subjects with known CAD
    - Number of subjects without known CAD
    - If applicable, enter the definition of known CAD
- Study Completion
  - Total population

- Number of subjects who received all study-specified diagnostic tests
- Number of subjects who completed follow-up phase to completion
- Female
  - Number of subjects who received all study-specified diagnostic tests
  - Number of subjects who completed follow-up phase to completion
- Male
  - Number of subjects who received all study-specified diagnostic tests
  - Number of subjects who completed follow-up phase to completion

## II. Baseline Demographics

- No known CAD reported
  - Age in years (Total, Female, and Male)
    - p value (Female vs. Male data)
    - Mean
    - SD
    - Min age
    - Max age
    - 25% IQR
    - 75% IQR
  - Ethnicity (Total, Female, and Male)
    - Hispanic or Latino
    - Not Hispanic or Latino
  - Race (Total, Female, and Male)
    - American Indian or Alaska Native
    - Asian
    - Black or African American
    - Native Hawaiian or other Pacific Islander
    - White
    - Other
    - Multiracial
    - Not reported
  - Was body size reported?
    - If yes, describe the measurement type and units.
    - If yes, provide the characteristics as reported (e.g. range, mean with standard deviation, etc.)
  - Was heart size reported?
    - If yes, describe the measurement type and units.
    - If yes, provide the characteristics as reported (e.g. range, mean with standard deviation, etc.)
  - Was functional status (exercise capacity) reported?
    - If yes, describe the measurement type and units
    - If yes, provide the characteristics as reported (e.g. range, mean with standard deviation, etc.)
- Mixed known/no known CAD reported
  - Age in years (Total, Female, and Male)
    - p value (Female vs. Male data)

- Mean
  - SD
  - Min age
  - Max age
  - 25% IQR
  - 75% IQR
- Ethnicity (Total, Female, and Male)
  - Hispanic or Latino
  - Not Hispanic or Latino
- Race (Total, Female, and Male)
  - American Indian or Alaska Native
  - Asian
  - Black or African American
  - Native Hawaiian or other Pacific Islander
  - White
  - Other
  - Multiracial
  - Not reported
- Was body size reported?
  - If yes, describe the measurement type and units.
  - If yes, provide the characteristics as reported (e.g. range, mean with standard deviation, etc.)
- Was heart size reported?
  - If yes, describe the measurement type and units.
  - If yes, provide the characteristics as reported (e.g. range, mean with standard deviation, etc.)
- Was functional status (exercise capacity) reported?
  - If yes, describe the measurement type and units
  - If yes, provide the characteristics as reported (e.g. range, mean with standard deviation, etc.)

### **III. NIT and Comparator Characteristics**

- NITs included in study
  - Electrocardiogram (exercise/stress or resting) = ECG
  - Echocardiography (with or without contrast) = ECHO
  - Exercise/stress radionuclide myocardial perfusion imaging (MPI). Includes:
    - Single proton emission computed tomography = SPECT,
    - Positron emission tomography = PET,
    - Scintigraphy
  - Cardiac perfusion and stress magnetic resonance imaging = CMR
  - Multidetector cardiac computed tomography angiography = CTA

### **IV. ECG Module**

- Type (multiple selections)
  - Exercise stress
  - Pharmacologic stress

- Multifunction Cardiogram [MCG]
- If exercise stress, type of exercise performed (multiple selections)
  - Treadmill
  - Bicycle
  - Step
  - Other (specify)
  - Not exercise stress
- If pharmacologic stress, type of agent used.
  - Dobutamine
  - Adenosine
  - Dipyridamole
  - Other (specify)
  - Not pharmacologic stress
- Definition of a positive result
  - $\geq 1$  mm ST depression
  - 1-2 mm ST depression
  - $\geq 2$  mm ST depression
  - Other (specify)

#### V. **Stress ECHO Module**

- Type of stressor (multiple selections)
  - Exercise stress
  - Pharmacologic stress
- If exercise stress, type of exercise performed (multiple selections)
  - Treadmill
  - Bicycle
  - Other (specify)
  - Not exercise stress
- If pharmacologic stress, type of agent used.
  - Dobutamine
  - Adenosine
  - Dipyridamole
  - Other (specify)
  - Not pharmacologic stress
- Was contrast agent used?
- Definition of a positive result
  - Wall Motion Abnormality (WMA) at rest and at stress
  - Wall Motion Abnormality (WMA) at stress, but not at rest
  - Wall Motion Abnormality (WMA) not otherwise specified
- If ECG used, definition of a positive result
  - $\geq 1$  mm ST depression
  - 1-2 mm ST depression
  - $\geq 2$  mm ST depression
  - Other (specify)

## **VI. Exercise/stress myocardial perfusion imaging Module**

- Modality
  - Single selection: SPECT, PET, scintigraphy
- Type of stressor
  - Exercise stress, Pharmacologic stress
  - If exercise stress, type of exercise performed.
    - Treadmill
    - Bicycle
    - Other (specify)
    - Not exercise stress
- If pharmacologic stress, type of agent used.
  - Dobutamine
  - Adenosine
  - Dipyridamole
  - Other (specify)
  - Not pharmacologic stress
- Radionuclide used
  - Technetium Tc 99m sestamibi (MIBI)
  - Thallous chloride TL-201 (thallium)
  - Fluorodeoxyglucose (FDG)
  - Other (specify)
- Definition of a positive result
  - Reported by Sum Stress Score (SSS)?
    - If yes, enter threshold value for positive result
  - Reported by % ischemic LV?
    - If yes, enter threshold value for positive result
  - Reported by evidence of ischemia in any segment?
    - If yes, enter number of segments considered a positive result
  - Reported by Transient Ischemic Dilation (or Total Perfusion Deficit)?
    - If yes, enter threshold value for positive result
- If ECG used, definition of a positive result
  - $\geq 1$  mm ST depression
  - 1-2 mm ST depression
  - $\geq 2$  mm ST depression
  - Other (specify)

## **VII. Cardiac perfusion and stress magnetic resonance imaging (CMR) Module**

- Type of test (multiple selections):
  - Dobutamine cine CMR
  - Vasodilator stress perfusion
  - Delayed enhanced (DE-CMR)
  - Other (specify)
- Type of stressor (multiple selections):
  - Exercise stress
  - Pharmacologic stress

- If exercise stress, type of exercise performed. If not exercise, select “Not exercise stress.” (multiple selections):
  - Bicycle
  - Other (specify)
  - Not exercise stress
- If pharmacologic stress, type of agent used (multiple selections):
  - Dobutamine
  - Adenosine
  - Dipyridamole
  - Other (specify w/free text field)
  - Not pharmacologic stress
- Was contrast agent used?
  - If yes, specify the contrast agent.
- Definition of positive result
  - Reported by perfusion defect?
    - If yes, enter threshold value for positive result
  - MRA of coronary arteries performed?
    - If yes, enter threshold value for positive result
  - Wall Motion Abnormalities (WMA) assessed?
    - If yes, select definition of positive result
      - Wall Motion Abnormality (WMA) at rest and at stress
      - Wall Motion Abnormality (WMA) at stress, but not at rest
      - Wall Motion Abnormality (WMA) not otherwise specified

### **VIII. Multidetector cardiac computed tomography angiography (CTA) Module**

- Was contrast agent used?
  - If yes, specify the contrast agent and dose (including units).
- Number of slices (multiple selections):
  - 4-slice
  - 16-slice
  - 32-slice
  - Other number less than 64 (specify w/text box)
  - 64-slice or greater
- Was calcium score testing performed?
- Definition of positive result (multiple selections):
  - $\geq 50\%$  stenosis
  - $\geq 70\%$  stenosis
  - $\geq 50\%$  Left Main
  - Other (specify)

### **IX. Diagnostic catheterization Module**

- Angiographic definition of disease (multiple selections):
  - $\geq 50\%$  stenosis

- $\geq 70\%$  stenosis
- $\geq 50\%$  Left Main
- Other (specify)

**X. Modality comparisons**

- Specify modality comparisons [one NIT to another (different) NIT, or a NIT to diagnostic cardiac catheterization].

**XI. Applicability to Key Questions**

- KQ 1:
  - What is the accuracy of one noninvasive technology (NIT) in diagnosing obstructive and nonobstructive CAD when compared to another NIT or to coronary angiography in women with chest pain syndrome?
    - Exercise electrocardiogram (ECG) stress test (including resting ECG technology, such as a multifunctional cardiogram)
    - Exercise/stress echocardiography (ECHO) with or without a contrast agent
    - Exercise/stress radionuclide myocardial perfusion imaging (including single proton emission computed tomography [SPECT] and positron emission tomography [PET])
    - Cardiac perfusion and stress magnetic resonance imaging (CMR)
    - Multidetector cardiac computed tomography angiography (CTA)
- KQ 2:
  - What are the predictors of diagnostic accuracy (age, race/ethnicity, body size, heart size, menopausal status, functional status, stress modality) of different NITs in women?
- KQ 3:
  - Is there evidence that the use of NITs (when compared to other NITs or to diagnostic cardiac catheterization) in women improves:
    - a. Risk stratification/prognostic information?
    - b. Decisionmaking regarding treatment options (e.g., revascularization, optimal medical therapy)?
    - c. Clinical outcomes (e.g., death, myocardial infarction, unstable angina, hospitalization, revascularization, angina relief, quality of life)?
- KQ 4:
  - Are there significant safety concerns/risks (i.e., radiation exposure, access site complications, contrast agent-induced nephropathy, nephrogenic systemic fibrosis, anaphylaxis, arrhythmias) associated with the use of different NITs to diagnose CAD in women with chest pain syndromes?

**XII. KQ 1**

- Which modality-level comparison(s) from the “NIT and Comparators” form does this data correspond to? Select all that apply. Space is provided for up to 6 modality-level comparisons.



- Enter total N for each category, NR for not reported, or NA for not applicable (Total population, Female, Male)
- Number of subjects with known CAD
- Number of subjects without known CAD
- If applicable, enter the definition of known CAD
- For the modality-level comparison(s) indicated above, does the article present data separately for both “no known CAD” and “mixed known/ no known CAD” populations?
  - If data is reported for a “no known CAD” population, enter age, ethnicity, race, body size, heart size, and functional status information for the “no known CAD” population in the section below labeled “No known CAD.”
  - If data is reported for a “mixed known/ no known CAD” population, enter age, ethnicity, race, body size, heart size, and functional status information for the mixed population in the section below labeled “Mixed known/ no known CAD.”
- Enter by Total population, Female, Male:
  - SD
  - Min age
  - Max age
  - 25% IQR
  - 75% IQR
  - Ethnicity
    - Hispanic
    - Non-Hispanic or Latino
  - Race
    - American Indian or Alaska Native
    - Asian
    - Black or African American
    - Native Hawaiian or other Pacific Islander
    - White
    - Other
    - Multiracial
    - Not reported
  - Was body size reported?
  - Was heart size reported?
  - Was functional status (exercise capacity) reported?

### **XIII. KQ 2**

- Indicate test modalities compared
- Indicate which predictive factor this data addresses
  - Age
  - Race/ethnicity
  - Body size
  - Heart size
  - Menopausal status
  - Functional status

- Stress modality
- Indicate CAD status of the population
  - No known CAD
  - Mixed known/ no known CAD
- Specify the female subgroups analyzed for this predictor and pair of test modalities. Columns are provided to capture up to 5 subgroup categories. Complete only the number needed to capture the data presented in the study.
- Define the groups
- Number of patients who received the index test
- Number of patients who received diagnostic cardiac cath
- Number of patients with adequate exercise for index test (if exercise is applicable)
- Number of patients with positive index test
- Number of patients with negative index test
- Disease prevalence (# of patients)
- Disease prevalence (%)
- True positive (# of patients)
- True negative (# of patients)
- False positive (# of patients)
- False negative (# of patients)
- Indeterminate or technically inadequate results (# of patients)
- Sensitivity (%)
- Sensitivity (Std dev)
- Sensitivity (Upper confidence interval bound)
- Sensitivity (Lower confidence interval bound)
- Specificity (%)
- Specificity (Std dev)
- Specificity (Upper confidence interval bound)
- Specificity (Lower confidence interval bound)
- Positive predictive value (%)
- Positive predictive value (Std dev)
- Positive predictive value (Upper confidence interval bound)
- Positive predictive value (Lower confidence interval bound)
- Negative predictive value (%)
- Negative predictive value (Std dev)
- Negative predictive value (Upper confidence interval bound)
- Negative predictive value (Lower confidence interval bound)
- Negative likelihood ratio
- Positive likelihood ratio
- Cath results
  - Number of patients with single-vessel disease
  - Number of patients with 2-vessel disease
  - Number of patients with 3-vessel disease
  - Number of patients with Left Main disease

#### **XIV. KQ 3**

- Indicate test modalities compared

- Indicate CAD status of the population
- Does the study provide data on risk stratification/prognostic information?
- Describe the risk/prognostic findings by gender
- Risk/prognostic information
  - Decisionmaking about treatment (Treatments may include: None, Medical management, Invasive management)
  - Describe any decisionmaking findings not captured by gender
- Clinical outcomes measured by Total population, Female, Male: (multiple selections):
  - MI
  - Unstable angina
  - Hospitalization
  - Mortality
  - Revascularization
  - Angina relief
  - Quality of life
  - Composite (specify)

**XV. KQ 4a**

- Indicate test modalities compared
- Indicate CAD status of the population
- Data by gender for each category of adverse outcome.
  - Average total body radiation exposure (specify units)
  - Access site complications
  - IV site complications
  - Contrast-agent induced nephropathy
  - Nephrogenic systemic fibrosis
  - Anaphylaxis
  - Arrhythmias
- If access site complications were reported, describe how these complications were defined.
- Does the article report tissue-level radiation data?
- Describe tissue-level radiation findings

**XVI. KQ 4b**

- Does the article report harms data broken down by any demographic factors other than gender? (Gender data is to be captured in form KQ 4a.)
- Indicate test modalities compared
- Indicate CAD status of the population
- Specify the categories for this subgroup analysis. Columns are provided to capture up to 5 categories. Complete only the number needed to capture the data presented in the study. Define the categories, then complete the tables below with as much information as is provided in the study.
- Harms
  - Average total body radiation exposure (specify units)
  - Access site complications

- IV site complications
- Contrast-agent induced nephropathy
- Nephrogenic systemic fibrosis
- Anaphylaxis
- Arrhythmias
- If access site complications were reported, describe how these complications were defined.
- Does the article report tissue-level radiation data?
- Describe tissue-level radiation findings

## **XVII. Quality Assessment**

- QUADAS Tool for Quality Assessment of Studies of Diagnostic Accuracy
- Answer each of the 14 questions below. A user's guide explaining each question and how to score your responses is available in the 2003 QUADAS article here: <http://www.biomedcentral.com/1471-2288/3/25>
  1. Was the spectrum of patients representative of the patients who will receive the test in practice?
  2. Were selection criteria clearly described?
  3. Is the reference standard likely to correctly classify the target condition?
  4. Is the time period between reference standard and index test short enough to be reasonably sure that the target condition did not change between the two tests?
  5. Did the whole sample or a random selection of the sample, receive verification using a reference standard of diagnosis?
  6. Did patients receive the same reference standard regardless of the index test result?
  7. Was the reference standard independent of the index test (i.e. the index test did not form part of the reference standard)?
  8. Was the execution of the index test described in sufficient detail to permit replication of the test?
  9. Was the execution of the reference standard described in sufficient detail to permit its replication?
  10. Were the index test results interpreted without knowledge of the results of the reference standard?
  11. Were the reference standard results interpreted without knowledge of the results of the index test?
  12. Were the same clinical data available when test results were interpreted as would be available when the test is used in practice?
  13. Were uninterpretable/ intermediate test results reported?
  14. Were withdrawals from the study explained?
- Summary Judgment
  - Good (low risk of bias). No major features that risk biased results. RCTs are considered a high study design type, but studies that include consecutive patients representative of the intended sample for whom diagnostic uncertainty exists may also meet this standard. A “good” study avoids the multiple biases to which

medical test studies are subject (e.g., use of an inadequate reference standard, verification bias), and key study features are clearly described, including the comparison groups, measurement of outcomes, and the characteristics of patients who failed to have actual state (diagnosis or prognosis) verified.

- Fair. Susceptible to some bias, but flaws not sufficient to invalidate the results. The study does not meet all the criteria required for a rating of good quality, but no flaw is likely to cause major bias. The study may be missing information, making it difficult to assess limitations and potential problems.
  - Poor (high risk of bias). Significant flaws that imply biases of various types that may invalidate the results. The study has significant biases determined a priori to be major or “fatal” (i.e., likely to make the results either uninterpretable or invalid).
- If the study is rated as “Fair” or “Poor,” provide rationale for decision.

### **XVIII. Applicability Assessment**

- Use the PICOTS format to identify specific issues that may limit the applicability of the study as described in the draft Methods Guide for Medical Test Reviews. Indicate the most important limitations affecting applicability, if any, from the list below.
- Population (P)
  - Source of population not described
  - Study population poorly specified
  - Key characteristics not reported
- Intervention (I)
  - Version/instrumentation not specified
  - Training/quality control not described
  - Screening and diagnostic uses mixed
- Comparator (C)
  - Gold standard not applied
  - Correlational data only
- Outcome of use of the test (O)
  - Failure to test “normals” or subset with gold standard
  - Precision of estimates not provided
- Clinical outcomes from test results (O)
  - Populations and study designs heterogeneous with varied findings
  - Data not stratified or adjusted for key predictors
- Timing (T)
  - Sequence of use of other diagnostics unclear
  - Time from results to treatment not reported
  - Order of testing varies across subjects and was not randomly assigned
- Setting (S)
  - Resources available to providers for diagnosis and treatment of condition vary widely
  - Provider type/specialty varies across settings
  - Comparability of care in international settings unclear

## Appendix C. List of Included Studies

- Abramson BL, Ruddy TD, deKemp RA, et al. Stress perfusion/metabolism imaging: a pilot study for a potential new approach to the diagnosis of coronary disease in women. *J Nucl Cardiol* 2000;7(3):205-12. PMID: 10888390
- Agati L, Renzi M, Sciomer S, et al. Transesophageal dipyridamole echocardiography for diagnosis of coronary artery disease. *J Am Coll Cardiol* 1992;19(4):765-70. PMID: 1545071
- Barolsky SM, Gilbert CA, Faruqi A, et al. Differences in electrocardiographic response to exercise of women and men: a non-Bayesian factor. *Circulation* 1979;60(5):1021-7. PMID: 487535
- Bjornstad K, Aakhus S, Hatle L. Comparison of digital dipyridamole stress echocardiography and upright bicycle stress echocardiography for identification of coronary artery stenosis. *Cardiology* 1995;86(6):514-20. PMID: 7585764
- Bokhari S, Shahzad A, Bergmann SR. Superiority of exercise myocardial perfusion imaging compared with the exercise ECG in the diagnosis of coronary artery disease. *Coron Artery Dis* 2008;19(6):399-404. PMID: 18955833
- Burgi Wegmann B, Sutsch G, Rickli H, et al. Gender and noninvasive diagnosis of coronary artery disease in women and men. *J Womens Health (Larchmt)* 2003;12(1):51-9. PMID: 12639369
- Chae SC, Heo J, Iskandrian AS, et al. Identification of extensive coronary artery disease in women by exercise single-photon emission computed tomographic (SPECT) thallium imaging. *J Am Coll Cardiol* 1993;21(6):1305-11. PMID: 8473634
- Cin VG, Tartanoglu O, Duzenli A, et al. The use of basic clinical and exercise variables in postmenopausal women for the diagnosis of coronary artery disease. *Int Angiol* 2000;9(3):135-137. PMID: 2000264000
- Coelho-Filho OR, Seabra LF, Mongeon FP, et al. Stress Myocardial Perfusion Imaging by CMR Provides Strong Prognostic Value to Cardiac Events Regardless of Patient's Sex. *JACC Cardiovasc Imaging* 2011;4(8):850-61. PMID: 21835377
- DePasquale EE, Nody AC, DePuey EG, et al. Quantitative rotational thallium-201 tomography for identifying and localizing coronary artery disease. *Circulation* 1988;77(2):316-27. PMID: 3257422
- Dewey M, Rutsch W, Hamm B. Is there a gender difference in noninvasive coronary imaging? Multislice computed tomography for noninvasive detection of coronary stenoses. *BMC Cardiovasc Disord* 2008;8:2. PMID: 18230167
- Dewey M, Teige F, Schnapauff D, et al. Noninvasive detection of coronary artery stenoses with multislice computed tomography or magnetic resonance imaging. *Ann Intern Med* 2006;145(6):407-15. PMID: 16983128
- Dewey M, Vavere AL, Arbab-Zadeh A, et al. Patient characteristics as predictors of image quality and diagnostic accuracy of MDCT compared with conventional coronary angiography for detecting coronary artery stenoses: CORE-64 Multicenter International Trial. *AJR Am J Roentgenol* 2010;194(1):93-102. PMID: 20028910
- Dharampall AS, Rossi A, Papadopoulou SL, et al. Is there a difference in the diagnostic accuracy of computed tomography coronary angiography between women and men? *Coron Artery Dis* 2011;22(6):421-427. PMID: 2011455721
- Dionisopoulos PN, Collins JD, Smart SC, et al. The value of dobutamine stress echocardiography for the detection of coronary artery disease in women. *J Am Soc Echocardiogr* 1997;10(8):811-7. PMID: 9356945
- Dodi C, Cortigiani L, Masini M, et al. The incremental prognostic value of pharmacological stress echo over exercise electrocardiography in women with chest pain of unknown origin. *Eur Heart J* 2001;22(2):145-52. PMID: 11161916
- Doyle M, Fuisz A, Kortright E, et al. The impact of myocardial flow reserve on the detection of coronary artery disease by perfusion imaging methods: an NHLBI WISE study. *J Cardiovasc Magn Reson* 2003;5(3):475-85. PMID: 12882078
- Doyle M, Weinberg N, Pohost GM, et al. Prognostic Value of Global MR Myocardial Perfusion Imaging in Women With Suspected Myocardial Ischemia and No Obstructive Coronary Disease Results From the NHLBI-Sponsored WISE (Women's Ischemia Syndrome Evaluation) Study. *JACC Cardiovasc Imaging* 2010;3(10):1030-6. PMID: 20947048
- Elhendy A, Geleijnse ML, van Domburg RT, et al. Gender differences in the accuracy of dobutamine stress echocardiography for the diagnosis of coronary

- artery disease. *Am J Cardiol* 1997;80(11):1414-8. PMID: 9399713
- Elhendy A, Schinkel AF, Bax JJ, et al. Accuracy of stress Tc-99m tetrofosmin myocardial perfusion tomography for the diagnosis and localization of coronary artery disease in women. *J Nucl Cardiol* 2006;13(5):629-34. PMID: 16945742
- Elhendy A, van Domburg RT, Bax JJ, et al. Noninvasive diagnosis of coronary artery stenosis in women with limited exercise capacity: comparison of dobutamine stress echocardiography and 99mTc sestamibi single-photon emission CT. *Chest* 1998;114(4):1097-104. PMID: 9792583
- Emmett L, Iwanochko RM, Freeman MR, et al. Reversible regional wall motion abnormalities on exercise technetium-99m-gated cardiac single photon emission computed tomography predict high-grade angiographic stenoses. *J Am Coll Cardiol* 2002;39(6):991-8. PMID: 11897441
- Faisal AW, Abid AR, Azhar M. Exercise Tolerance Test: a comparison between true positive and false positive test results. *J Ayub Med Coll Abbottabad* 2007;19(4):71-4. PMID: 18693603
- Friedman TD, Greene AC, Iskandrian AS, et al. Exercise thallium-201 myocardial scintigraphy in women: correlation with coronary arteriography. *Am J Cardiol* 1982;49(7):1632-7. PMID: 7081050
- Gebker R, Jahnke C, Hucko T, et al. Dobutamine stress magnetic resonance imaging for the detection of coronary artery disease in women. *Heart* 2010;96(8):616-20. PMID: 19687013
- Gentile R, Vitarelli A, Schillaci O, et al. Diagnostic accuracy and prognostic implications of stress testing for coronary artery disease in the elderly. *Ital Heart J* 2001;2(7):539-45. PMID: 11501963
- Guiteras P, Chaitman BR, Waters DD, et al. Diagnostic accuracy of exercise ECG lead systems in clinical subsets of women. *Circulation* 1982;65(7):1465-74. PMID: 7074802
- Gulati M, Pratap P, Kansal P, et al. Gender differences in the value of ST-segment depression during adenosine stress testing. *Am J Cardiol* 2004;94(8):997-1002. PMID: 15476611
- Hlatky MA, Pryor DB, Harrell FE, Jr., et al. Factors affecting sensitivity and specificity of exercise electrocardiography. Multivariable analysis. *Am J Med* 1984;77(1):64-71. PMID: 6741986
- Ho YL, Wu CC, Huang PJ, et al. Assessment of coronary artery disease in women by dobutamine stress echocardiography: comparison with stress thallium-201 single-photon emission computed tomography and exercise electrocardiography. *Am Heart J* 1998;135(4):655-62. PMID: 9539482
- Hoiland-Carlsen PF, Johansen A, Christensen HW, et al. Usefulness of the exercise electrocardiogram in diagnosing ischemic or coronary heart disease in patients with chest pain. *Am J Cardiol* 2005;95(1):96-9. PMID: 15619400
- Hoiland-Carlsen PF, Johansen A, Vach W, et al. High probability of disease in angina pectoris patients: is clinical estimation reliable? *Can J Cardiol* 2007;23(8):641-7. PMID: 17593989
- Hosokawa J, Shen JT, Imhoff M. Computerized 2-lead resting ECG analysis for the detection of relevant coronary artery stenosis in comparison with angiographic findings. *Congest Heart Fail* 2008;14(5):251-60. PMID: 18983288
- Hung J, Chaitman BR, Lam J, et al. Noninvasive diagnostic test choices for the evaluation of coronary artery disease in women: a multivariate comparison of cardiac fluoroscopy, exercise electrocardiography and exercise thallium myocardial perfusion scintigraphy. *J Am Coll Cardiol* 1984;4(1):8-16. PMID: 6736458
- Jenkins SM, Johnston N, Hawkins NM, et al. Limited clinical utility of CT coronary angiography in a district hospital setting. *QJM* 2010. PMID: 20847015
- Johansen A, Hoiland-Carlsen PF, Christensen HW, et al. Observer variability in the evaluation of dual-isotope Tl-201/Tc-99m sestamibi rest/stress myocardial perfusion SPECT in men and women with known or suspected stable angina pectoris. *J Nucl Cardiol* 2004;11(6):710-8. PMID: 15592195
- Johnson BD, Shaw LJ, Buchthal SD, et al. Prognosis in women with myocardial ischemia in the absence of obstructive coronary disease: results from the National Institutes of Health-National Heart, Lung, and Blood Institute-Sponsored Women's Ischemia Syndrome Evaluation (WISE). *Circulation* 2004;109(24):2993-9. PMID: 15197152
- Kaminek M, Myslivecek M, Husak V, et al. The accuracy of myocardial perfusion SPECT imaging in the evaluation of coronary artery disease in women and men. *Nucl Med Rev Cent East Eur* 2001;4(2):69-72. PMID: 14600887
- Kiat H, Van Train KF, Maddahi J, et al. Development and prospective application of quantitative 2-day stress-rest Tc-99m methoxy isobutyl isonitrile SPECT for the diagnosis of

- coronary artery disease. *Am Heart J* 1990;120(6 Pt 1):1255-66. PMID: 2248175
- Klem I, Greulich S, Heitner JF, et al. Value of cardiovascular magnetic resonance stress perfusion testing for the detection of coronary artery disease in women. *JACC Cardiovasc Imaging* 2008;1(4):436-45. PMID: 19356464
- Koide Y, Yotsukura M, Yoshino H, et al. A new coronary artery disease index of treadmill exercise electrocardiograms based on the step-up diagnostic method. *Am J Cardiol* 2001;87(2):142-7. PMID: 11152828
- Langer C, Peterschroder A, Franzke K, et al. Noninvasive coronary angiography focusing on calcification: multislice computed tomography compared with magnetic resonance imaging. *J Comput Assist Tomogr* 2009;33(2):179-85. PMID: 19346842
- Laurienzo JM, Cannon RO, 3rd, Quyyumi AA, et al. Improved specificity of transesophageal dobutamine stress echocardiography compared to standard tests for evaluation of coronary artery disease in women presenting with chest pain. *Am J Cardiol* 1997;80(11):1402-7. PMID: 9399711
- Lehmkuhl HB, Siniawski H, Lehmkuhl E, et al. Value and limitations of dobutamine stress echocardiography in women with suspected coronary artery disease. *Zeitschrift fur Herz-, Thorax- und Gefasschirurgie* 2007;21(6):250-258. PMID: 2008012568
- Lewandowski M, Szwed H, Kowalik I. Searching for the optimal strategy for the diagnosis of stable coronary artery disease. Cost-effectiveness of the new algorithm. *Cardiol J* 2007;14(6):544-51. PMID: 18651520
- Lewis JF, Lin L, McGorray S, et al. Dobutamine stress echocardiography in women with chest pain. Pilot phase data from the National Heart, Lung and Blood Institute Women's Ischemia Syndrome Evaluation (WISE). *J Am Coll Cardiol* 1999;33(6):1462-8. PMID: 10334409
- Lewis JF, McGorray S, Lin L, et al. Exercise treadmill testing using a modified exercise protocol in women with suspected myocardial ischemia: findings from the National Heart, Lung and Blood Institute-sponsored Women's Ischemia Syndrome Evaluation (WISE). *Am Heart J* 2005;149(3):527-33. PMID: 15864243
- Lu C, Lu F, Fragasso G, et al. Comparison of exercise electrocardiography, technetium-99m sestamibi single photon emission computed tomography, and dobutamine and dipyridamole echocardiography for detection of coronary artery disease in hypertensive women. *Am J Cardiol* 2010;105(9):1254-60. PMID: 20403475
- Maffei E, Seitun S, Martini C, et al. CT coronary angiography and exercise ECG in a population with chest pain and low-to-intermediate pre-test likelihood of coronary artery disease. *Heart* 2010;96(24):1973-9. PMID: 21051457
- Majstorov V, Pop Gjorceva D, Vaskova O, et al. Gender differences in detecting coronary artery disease with dipyridamole stress myocardial perfusion imaging using 99m-Tc sestamibi gated SPECT. *Prilozi* 2005;26(1):93-102. PMID: 16118618
- Mak KH, Ang ES, Goh AS, et al. Myocardial perfusion imaging with technetium-99m sestamibi SPECT in the evaluation of coronary artery disease. *Australas Radiol* 1995;39(2):112-7. PMID: 7605313
- Marwick T, D'Hondt AM, Baudhuin T, et al. Optimal use of dobutamine stress for the detection and evaluation of coronary artery disease: combination with echocardiography or scintigraphy, or both? *J Am Coll Cardiol* 1993;22(1):159-67. PMID: 8509537
- Marwick TH, Anderson T, Williams MJ, et al. Exercise echocardiography is an accurate and cost-efficient technique for detection of coronary artery disease in women. *J Am Coll Cardiol* 1995;26(2):335-41. PMID: 7608432
- Masini M, Picano E, Lattanzi F, et al. High dose dipyridamole-echocardiography test in women: correlation with exercise-electrocardiography test and coronary arteriography. *J Am Coll Cardiol* 1988;12(3):682-5. PMID: 3403825
- Mazeika P, Nihoyannopoulos P, Joshi J, et al. Uses and limitations of high dose dipyridamole stress echocardiography for evaluation of coronary artery disease. *Br Heart J* 1992;67(2):144-9. PMID: 1540434
- Meijboom WB, Meijs MF, Schuijf JD, et al. Diagnostic accuracy of 64-slice computed tomography coronary angiography: a prospective, multicenter, multivendor study. *J Am Coll Cardiol* 2008;52(25):2135-44. PMID: 19095130
- Meijboom WB, Weustink AC, Pugliese F, et al. Comparison of diagnostic accuracy of 64-slice computed tomography coronary angiography in women versus men with angina pectoris. *Am J Cardiol* 2007;100(10):1532-7. PMID: 17996514



- Merkle N, Wöhrle J, Nusser T, et al. Diagnostic performance of magnetic resonance first pass perfusion imaging is equally potent in female compared to male patients with coronary artery disease. *Clin Res Cardiol* 2010;99(1):21-8. PMID: 19756814
- Merz CN, Kelsey SF, Pepine CJ, et al. The Women's Ischemia Syndrome Evaluation (WISE) study: protocol design, methodology and feasibility report. *J Am Coll Cardiol* 1999;33(6):1453-61. PMID: 10334408
- Michaelides AP, Fourlas CA, Chatzistamatiou EI, et al. QRS score improves diagnostic ability of treadmill exercise testing in women. *Coron Artery Dis* 2007;18(4):313-8. PMID: 17496496
- Mieres JH, Makaryus AN, Cacciabauda JM, et al. Value of electrocardiographically gated single-photon emission computed tomographic myocardial perfusion scintigraphy in a cohort of symptomatic postmenopausal women. *Am J Cardiol* 2007;99(8):1096-9. PMID: 17437734
- Mieres JH, Shaw LJ, Hendel RC, et al. The WOMEN study: what is the optimal method for ischemia evaluation in women? A multi-center, prospective, randomized study to establish the optimal method for detection of coronary artery disease (CAD) risk in women at an intermediate-high pretest likelihood of CAD: study design. *J Nucl Cardiol* 2009;16(1):105-12. PMID: 19152135
- Miller JM, Dewey M, Vavere AL, et al. Coronary CT angiography using 64 detector rows: methods and design of the multi-centre trial CORE-64. *Eur Radiol* 2009;19(4):816-28. PMID: 18998142
- Miller TD, Roger VL, Milavetz JJ, et al. Assessment of the exercise electrocardiogram in women versus men using tomographic myocardial perfusion imaging as the reference standard. *Am J Cardiol* 2001;87(7):868-73. PMID: 11274942
- Mohiuddin SM, Ravage CK, Esterbrooks DJ, et al. The comparative safety and diagnostic accuracy of adenosine myocardial perfusion imaging in women versus men. *Pharmacotherapy* 1996;16(4):646-51. PMID: 8840371
- Morise AP. Are the American College of Cardiology/American Heart Association guidelines for exercise testing for suspected coronary artery disease correct? *Chest* 2000;118(2):535-41. PMID: 10936152
- Morise AP, Diamond GA. Comparison of the sensitivity and specificity of exercise electrocardiography in biased and unbiased populations of men and women. *Am Heart J* 1995-A;130(4):741-7. PMID: 7572581
- Morise AP, Diamond GA, Detrano R, et al. Incremental value of exercise electrocardiography and thallium-201 testing in men and women for the presence and extent of coronary artery disease. *Am Heart J* 1995-B;130(2):267-76. PMID: 7631606
- Morise AP, Lauer MS, Froelicher VF. Development and validation of a simple exercise test score for use in women with symptoms of suspected coronary artery disease. *Am Heart J* 2002;144(5):818-25. PMID: 12422150
- Morise AP, Olson MB, Merz CN, et al. Validation of the accuracy of pretest and exercise test scores in women with a low prevalence of coronary disease: the NHLBI-sponsored Women's Ischemia Syndrome Evaluation (WISE) study. *Am Heart J* 2004;147(6):1085-92. PMID: 15199360
- Nascimento BR, Chequer G, Barbosa MP, et al. Stable angina and ST-segment depression: diagnostic value in patients undergoing coronary angiography. *Rev Port Cardiol* 2008;27(12):1567-80. PMID: 19280997
- Ozdemir K, Altunkeser BB, Aydin M, et al. New parameters in the interpretation of exercise testing in women: QTc dispersion and QT dispersion ratio difference. *Clin Cardiol* 2002;25(4):187-92. PMID: 12000077
- Pundziute G, Schuijf JD, Jukema JW, et al. Gender influence on the diagnostic accuracy of 64-slice multislice computed tomography coronary angiography for detection of obstructive coronary artery disease. *Heart* 2008;94(1):48-52. PMID: 17540687
- Raman SV, Donnally MR, McCarthy B. Dobutamine stress cardiac magnetic resonance imaging to detect myocardial ischemia in women. *Prev Cardiol* 2008;11(3):135-40. PMID: 18607148
- Richardson MT, Holly RG, Amsterdam EA, et al. The value of ten common exercise tolerance test measures in predicting coronary disease in symptomatic females. *Cardiology* 1995;86(3):243-8. PMID: 7614498
- Robert AR, Melin JA, Detry JM. Logistic discriminant analysis improves diagnostic accuracy of exercise testing for coronary artery disease in women. *Circulation* 1991;83(4):1202-9. PMID: 2013142
- Roger VL, Pellikka PA, Bell MR, et al. Sex and test verification bias. Impact on the diagnostic value of

- exercise echocardiography. *Circulation* 1997;95(2):405-10. PMID: 9008457
- Rollan MJ, San Roman JA, Vilacosta I, et al. Dobutamine stress echocardiography in the diagnosis of coronary artery disease in women with chest pain: comparison with different noninvasive tests. *Clin Cardiol* 2002;25(12):559-64. PMID: 12492125
- San Roman JA, Vilacosta I, Castillo JA, et al. Selection of the optimal stress test for the diagnosis of coronary artery disease. *Heart* 1998;80(4):370-6. PMID: 9875115
- Sanfilippo AJ, Abdollah H, Knott TC, et al. Stress echocardiography in the evaluation of women presenting with chest pain syndrome: a randomized, prospective comparison with electrocardiographic stress testing. *Can J Cardiol* 2005;21(5):405-12. PMID: 15861257
- Santana-Boado C, Candell-Riera J, Castell-Conesa J, et al. Diagnostic accuracy of technetium-99m-MIBI myocardial SPECT in women and men. *J Nucl Med* 1998;39(5):751-5. PMID: 9591568
- Sawada SG, Ryan T, Fineberg NS, et al. Exercise echocardiographic detection of coronary artery disease in women. *J Am Coll Cardiol* 1989;14(6):1440-7. PMID: 2809000
- Schupbach WM, Emese B, Loretan P, et al. Non-invasive diagnosis of coronary artery disease using cardiogoniometry performed at rest. *Swiss Med Wkly* 2008;138(15-16):230-8. PMID: 18431698
- Severi S, Picano E, Michelassi C, et al. Diagnostic and prognostic value of dipyridamole echocardiography in patients with suspected coronary artery disease. Comparison with exercise electrocardiography. *Circulation* 1994;89(3):1160-73. PMID: 8124803
- Sharir T, Germano G, Waechter PB, et al. A new algorithm for the quantitation of myocardial perfusion SPECT. II: validation and diagnostic yield. *J Nucl Med* 2000;41(4):720-7. PMID: 10768575
- Shaw LJ, Mieres JH, Hendel RH, et al. Comparative effectiveness of exercise electrocardiography with or without myocardial perfusion single photon emission computed tomography in women with suspected coronary artery disease: results From the What Is the Optimal Method for Ischemia Evaluation in Women (WOMEN) Trial. *Circulation* 2011. PMID: 21844080
- Shi H, Santana CA, Rivero A, et al. Normal values and prospective validation of transient ischaemic dilation index in 82Rb PET myocardial perfusion imaging. *Nucl Med Commun* 2007;28(11):859-63. PMID: 17901769
- Shin JH, Shiota T, Kim YJ, et al. False-positive exercise echocardiograms: impact of sex and blood pressure response. *Am Heart J* 2003;146(5):914-9. PMID: 14597944
- Shivalkar B, Goovaerts I, Salgado RA, et al. Multislice cardiac computed tomography in symptomatic middle-aged women. *Ann Med* 2007;39(4):290-7. PMID: 17558600
- Siegler JC, Rehman S, Bhumireddy GP, et al. The Accuracy of the Electrocardiogram during Exercise Stress Test Based on Heart Size. *PLoS One* 2011;6(8):e23044. PMID: 21857990
- Sinha DP, Das M, Banerjee AK, et al. Comparative study to assess whether high sensitive C-reactive protein and carotid intima media thickness improve the predictive accuracy of exercise stress testing for coronary artery disease in perimenopausal women with typical angina. *J Indian Med Assoc* 2008;106(2):86, 88, 90 passim. PMID: 18705250
- Sketch MH, Mohiuddin SM, Lynch JD, et al. Significant sex differences in the correlation of electrocardiographic exercise testing and coronary arteriograms. *Am J Cardiol* 1975;36(2):169-73. PMID: 1155337
- Slavich GA, Guerra UP, Morocutti G, et al. Feasibility of simultaneous Tc99m sestamibi and 2D-echo cardiac imaging during dobutamine pharmacologic stress. Preliminary results in a female population. *Int J Card Imaging* 1996;12(2):113-8. PMID: 8864790
- Slomka PJ, Nishina H, Abidov A, et al. Combined quantitative supine-prone myocardial perfusion SPECT improves detection of coronary artery disease and normalcy rates in women. *J Nucl Cardiol* 2007;14(1):44-52. PMID: 17276305
- Slomka PJ, Nishina H, Berman DS, et al. Automated quantification of myocardial perfusion SPECT using simplified normal limits. *J Nucl Cardiol* 2005;12(1):66-77. PMID: 15682367
- Svart K, Lehtinen R, Nieminen T, et al. Exercise electrocardiography detection of coronary artery disease by. *International Journal of Cardiology* 2010;140(2):182-188. PMID: 2010202240
- Taillefer R, DePuey EG, Udelson JE, et al. Comparative diagnostic accuracy of Tl-201 and Tc-99m sestamibi SPECT imaging (perfusion and ECG-gated SPECT) in detecting coronary artery disease in

women. *J Am Coll Cardiol* 1997;29(1):69-77. PMID: 8996297

Takeuchi M, Sonoda S, Miura Y,, et al. Comparative diagnostic value of dobutamine stress echocardiography and stress thallium-201 single-photon-emission computed tomography for detecting coronary artery disease in women. *Coron Artery Dis* 1996;7(11):831-5. PMID: 8993941

Travin MI, Katz MS, Moulton AW,, et al. Accuracy of dipyridamole SPECT imaging in identifying individual coronary stenoses and multivessel disease in women versus men. *J Nucl Cardiol* 2000;7(3):213-20. PMID: 10888391

Van Train KF, Garcia EV, Maddahi J,, et al. Multicenter trial validation for quantitative analysis of same-day rest-stress technetium-99m-sestamibi myocardial tomograms. *J Nucl Med* 1994;35(4):609-18. PMID: 8151383

Vashist A, Collins D, Prasad Y,, et al. Does cardiac SPECT using attenuation and scatter correction accurately predict coronary artery disease in a minority women population? *Med Sci Monit* 2007;13(9):CR386-390. PMID: 17767117

Weiner DA. Accuracy of cardiokymography during exercise testing: results of a multicenter study. *J Am Coll Cardiol* 1985;6(3):502-9. PMID: 3897339

Weiner DA, Ryan TJ, McCabe CH,, et al. Exercise stress testing. Correlations among history of angina, ST-segment response and prevalence of coronary-artery disease in the Coronary Artery Surgery Study (CASS). *N Engl J Med* 1979;301(5):230-5. PMID: 449990

Weustink AC, Meijboom WB, Mollet NR,, et al. Reliable high-speed coronary computed tomography in symptomatic patients. *J Am Coll Cardiol* 2007;50(8):786-94. PMID: 17707184

Weustink AC, Mollet NR, Neeffjes LA,, et al. Preserved diagnostic performance of dual-source CT coronary angiography with reduced radiation exposure and cancer risk. *Radiology* 2009;252(1):53-60. PMID: 19451542

Williams MJ, Marwick TH, O'Gorman D,, et al. Comparison of exercise echocardiography with an exercise score to diagnose coronary artery disease in women. *Am J Cardiol* 1994;74(5):435-8. PMID: 8059721

Wolak A, Slomka PJ, Fish MB,, et al. Quantitative diagnostic performance of myocardial perfusion SPECT with attenuation correction in women. *J Nucl Med* 2008;49(6):915-22. PMID: 18483092

Wong Y, Rodwell A, Dawkins S,, et al. Sex differences in investigation results and treatment in subjects referred for investigation of chest pain. *Heart* 2001;85(2):149-52. PMID: 11156662

Yamauchi K, Simonson E, Dahl JC,, et al. Sex differences in submaximal exercise tests correlation with coronary cineangiography in 133 patients. *Nagoya J Med Sci* 1985;47(1-2):67-75. PMID: 3990772

Yeih DF, Huang PJ, Ho YL. Enhanced diagnosis of coronary artery disease in women by dobutamine thallium-201 ST-segment/heart rate slope and thallium-201 myocardial SPECT. *J Formos Med Assoc* 2007;106(10):832-9. PMID: 17964962

## Appendix D. Quality and Applicability of Included Studies

**Table D-1. Quality, applicability, and relevant Key Questions (KQs)**

| Study                       | NIT Modality  | KQ 1 | KQ 2 | KQ 3 | KQ 4 | Quality | Limitations to Applicability  |
|-----------------------------|---|------|------|------|------|---------|---|
| Abramson, et al., 2000      | <ul style="list-style-type: none"> <li>• SPECT/PET/Scintigraphy</li> </ul>                | X    |      |      |      | Fair    | <ul style="list-style-type: none"> <li>• Key characteristics not reported</li> <li>• Failure to test “normals” or subset with gold standard</li> <li>• Data not stratified or adjusted for key predictors</li> </ul>  |
| Agati, et al., 1992         | <ul style="list-style-type: none"> <li>• ECG</li> <li>• ECHO</li> </ul>                   | X    |      |      |      | Fair    | <ul style="list-style-type: none"> <li>• Source of population not described</li> <li>• Key characteristics not reported</li> <li>• Training/quality control not described</li> <li>• Precision of estimates not provided</li> </ul>   |
| Barolsky, et al., 1979      | <ul style="list-style-type: none"> <li>• ECG</li> </ul>                                   | X    |      |      |      | Fair    | <ul style="list-style-type: none"> <li>• Key characteristics not reported</li> <li>• Training/quality control not described</li> <li>• Precision of estimates not provided</li> <li>• Data not stratified or adjusted for key predictors</li> </ul>   |
| Bjornstad, et al., 1995     | <ul style="list-style-type: none"> <li>• ECHO</li> </ul>                                  | X    |      |      |      | Fair    | <ul style="list-style-type: none"> <li>• Key characteristics not reported</li> <li>• Training/quality control not described</li> <li>• Failure to test “normals” or subset with gold standard</li> <li>• Precision of estimates not provided</li> </ul>   |
| Bokhari, et al., 2008       | <ul style="list-style-type: none"> <li>• ECG</li> <li>• SPECT/PET/scintigraphy</li> </ul> | X    |      |      |      | Poor    | <ul style="list-style-type: none"> <li>• Training/quality control not described</li> <li>• Data not stratified or adjusted for key predictors</li> <li>• Sequence of use of other diagnostics unclear</li> </ul>  |
| Burgi Wegmann, et al., 2003 | <ul style="list-style-type: none"> <li>• ECG</li> </ul>                                   | X    |      |      |      | Fair    | <ul style="list-style-type: none"> <li>• Source of population not described</li> <li>• Study population poorly specified</li> <li>• Version/instrumentation not specified</li> <li>• Training/quality control not described</li> <li>• Sequence of use of other diagnostics unclear</li> <li>• Comparability of care in international settings unclear</li> </ul> |
| Chae, et al., 1993          | <ul style="list-style-type: none"> <li>• SPECT/PET/scintigraphy</li> </ul>                | X    |      |      |      | Good    | <ul style="list-style-type: none"> <li>• Training/quality control not described</li> <li>• Precision of estimates not provided</li> </ul>   |
| Cin, et al., 2000           | <ul style="list-style-type: none"> <li>• ECG</li> </ul>                                   | X    | X    |      |      | Poor    | <ul style="list-style-type: none"> <li>• Training/quality control not described</li> <li>• Precision of estimates not provided</li> <li>• Data not stratified or adjusted for key predictors</li> </ul>   |

| Study                                | NIT Modality  | KQ 1 | KQ 2 | KQ 3 | KQ 4 | Quality | Limitations to Applicability   |
|--------------------------------------|---|------|------|------|------|---------|--|
| Coelho-Filho, et al., 2011           | <ul style="list-style-type: none"> <li>• CMR</li> <li>• Coronary CTA</li> </ul>           | X    |      | X    |      | Fair    | <ul style="list-style-type: none"> <li>• Training/quality control not described</li> <li>• Gold standard not applied</li> </ul>  |
| DePasquale, et al., 1988             | <ul style="list-style-type: none"> <li>• SPECT/PET/scintigraphy</li> </ul>                | X    |      |      |      | Poor    | <ul style="list-style-type: none"> <li>• Source of population not described</li> <li>• Study population poorly specified</li> <li>• Key characteristics not reported</li> <li>• Precision of estimates not provided</li> </ul>   |
| Dewey, et al., 2008 <sup>a</sup>     | <ul style="list-style-type: none"> <li>• Coronary CTA</li> </ul>                          | X    |      |      | X    | Good    | <ul style="list-style-type: none"> <li>• Precision of estimates not provided</li> </ul>  |
| Dewey, et al., 2010 <sup>b</sup>     | <ul style="list-style-type: none"> <li>• CTA</li> </ul>                                   | X    |      |      |      | Good    | <ul style="list-style-type: none"> <li>• Precision of estimates not provided</li> </ul>  |
| Dharampal, et al., 2011 <sup>c</sup> | <ul style="list-style-type: none"> <li>• Coronary CTA</li> </ul>                          | X    |      |      | X    | Fair    | <ul style="list-style-type: none"> <li>• Screening and diagnostic uses mixed</li> <li>• Comparability of care in international settings unclear</li> <li>• Limitation from use of two types of CTA technology within study</li> </ul>  |
| Dionisopoulos, et al., 1997          | <ul style="list-style-type: none"> <li>• ECHO</li> </ul>                                  | X    |      |      |      | Fair    | <ul style="list-style-type: none"> <li>• Study population poorly specified</li> <li>• Key characteristics not reported</li> <li>• Training/quality control not described</li> <li>• Precision of estimates not provided</li> </ul>   |
| Dodi, et al., 2001                   | <ul style="list-style-type: none"> <li>• ECG</li> <li>• ECHO</li> </ul>                   |      |      | X    |      | Good    | <ul style="list-style-type: none"> <li>• Training/quality control not described</li> <li>• Gold standard not applied</li> <li>• Order of testing varied across subjects and was not randomly assigned</li> </ul>   |
| Doyle, et al., 2003 <sup>d</sup>     | <ul style="list-style-type: none"> <li>• SPECT/PET/scintigraphy</li> <li>• CMR</li> </ul> | X    |      |      |      | Good    | <ul style="list-style-type: none"> <li>• Precision of estimates not provided</li> <li>• Order of testing varied across subjects and was not randomly assigned</li> </ul>   |
| Doyle, et al., 2010 <sup>d</sup>     | <ul style="list-style-type: none"> <li>• CMR</li> </ul>                                   |      |      | X    |      | Fair    | <ul style="list-style-type: none"> <li>• Source of population not described</li> <li>• Training/quality control not described</li> <li>• Precision of estimates not provided</li> <li>• Resources available to providers for diagnosis and treatment of condition varied widely</li> </ul> |
| Elhendy, et al., 1997                | <ul style="list-style-type: none"> <li>• ECH</li> <li>• ECHO</li> </ul>                   | X    |      |      | X    | Fair    | <ul style="list-style-type: none"> <li>• Source of population not described</li> <li>• Comparability of care in international settings unclear</li> </ul>  |

| Study                  | NIT Modality   | KQ 1 | KQ 2 | KQ 3 | KQ 4 | Quality | Limitations to Applicability  |
|------------------------|--|------|------|------|------|---------|---|
| Elhendy, et al., 2006  | <ul style="list-style-type: none"> <li>• SPECT/PET/scintigraphy</li> </ul>                 | X    |      |      |      | Fair    | <ul style="list-style-type: none"> <li>• Study population poorly specified</li> <li>• Key characteristics not reported</li> <li>• Training/quality control not described</li> <li>• Failure to test “normals” or subset with gold standard</li> <li>• Comparability of care in international settings unclear</li> </ul>  |
| Elhendy, et al., 1998  | <ul style="list-style-type: none"> <li>• ECHO</li> <li>• SPECT/PET/scintigraphy</li> </ul> | X    |      |      |      | Good    | <ul style="list-style-type: none"> <li>• Training/quality control not described</li> </ul>  |
| Emmett, et al., 2002   | <ul style="list-style-type: none"> <li>• SPECT/PET/scintigraphy</li> </ul>                 | X    |      |      |      | Fair    | <ul style="list-style-type: none"> <li>• Source of population not described</li> <li>• Key characteristics not reported</li> <li>• Precision of estimates not provided</li> <li>• Comparability of care in international settings unclear</li> </ul>  |
| Faisal, et al., 2007   | <ul style="list-style-type: none"> <li>• ECG</li> </ul>                                    | X    |      |      |      | Poor    | <ul style="list-style-type: none"> <li>• Source of population not described</li> <li>• Study population poorly specified</li> <li>• Training/quality control not described</li> <li>• Failure to test “normals” or subset with gold standard</li> <li>• Precision of estimates not provided</li> <li>• Sequence of use of other diagnostics unclear</li> <li>• Resources available to providers for diagnosis and treatment of condition varied widely</li> </ul> |
| Friedman, et al., 1982 | <ul style="list-style-type: none"> <li>• ECG</li> <li>• SPECT/PET/scintigraphy</li> </ul>  | X    |      |      |      | Fair    | <ul style="list-style-type: none"> <li>• Source of population not described</li> <li>• Precision of estimates not provided</li> </ul>   |
| Gebker, et al., 2010   | <ul style="list-style-type: none"> <li>• CMR</li> </ul>                                    | X    | X    |      | X    | Good    | <ul style="list-style-type: none"> <li>• Source of population not described</li> <li>• Precision of estimates not provided</li> </ul>   |
| Gentile, et al., 2001  | <ul style="list-style-type: none"> <li>• ECG</li> <li>• SPECT/PET/scintigraphy</li> </ul>  | X    |      |      |      | Good    | <ul style="list-style-type: none"> <li>• Training/quality control not described</li> <li>• Data not stratified or adjusted for key predictors</li> </ul>  |
| Guiteras, et al., 1982 | <ul style="list-style-type: none"> <li>• ECG</li> </ul>                                    | X    |      |      |      | Fair    | <ul style="list-style-type: none"> <li>• Source of population not described</li> <li>• Training/quality control not described</li> <li>• Precision of estimates not provided</li> <li>• Comparability of care in international settings unclear</li> </ul>  |
| Gulati, et al., 2004   | <ul style="list-style-type: none"> <li>• SPECT/PET/scintigraphy</li> </ul>                 | X    |      |      |      | Fair    | <ul style="list-style-type: none"> <li>• Source of population not described</li> <li>• Comparability of care in international settings unclear</li> </ul>   |

| Study                              | NIT Modality   | KQ 1 | KQ 2 | KQ 3 | KQ 4 | Quality | Limitations to Applicability   |
|------------------------------------|--|------|------|------|------|---------|--|
| Hlatky, et al., 1984               | <ul style="list-style-type: none"> <li>• ECG</li> </ul>                                    | X    |      |      |      | Poor    | <ul style="list-style-type: none"> <li>• Source of population not described</li> <li>• Key characteristics not reported</li> <li>• Version/instrumentation not specified</li> <li>• Training/quality control not described</li> <li>• Precision of estimates not provided</li> <li>• Time from results to treatment not reported</li> </ul>              |
| Ho, et al., 1998                   | <ul style="list-style-type: none"> <li>• ECHO</li> <li>• SPECT/PET/scintigraphy</li> </ul> | X    | X    |      | X    | Fair    | <ul style="list-style-type: none"> <li>• Source of population not described</li> <li>• Precision of estimates not provided</li> <li>• Comparability of care in international settings unclear</li> </ul>   |
| Hoiland-Carlsen, et al., 2005      | <ul style="list-style-type: none"> <li>• ECG</li> <li>• SPECT/PET/scintigraphy</li> </ul>  | X    |      |      |      | Fair    | <ul style="list-style-type: none"> <li>• Source of population not described</li> <li>• Study population poorly specified</li> <li>• Version/instrumentation not specified</li> <li>• Training/quality control not described</li> <li>• Precision of estimates not provided</li> <li>• Comparability of care in international settings unclear</li> </ul> |
| Hoiland-Carlsen, et al., 2007      | <ul style="list-style-type: none"> <li>• SPECT/PET/scintigraphy</li> </ul>                 | X    |      |      |      | Good    | <ul style="list-style-type: none"> <li>• Training/quality control not described</li> </ul>   |
| Hosokawa, et al., 2008             | <ul style="list-style-type: none"> <li>• ECG</li> </ul>                                    | X    |      |      |      | Fair    | <ul style="list-style-type: none"> <li>• Source of population not described</li> <li>• Study population poorly specified</li> <li>• Key characteristics not reported</li> <li>• Precision of estimates not provided</li> <li>• Time from results to treatment not reported</li> <li>• Comparability of care in international settings unclear</li> </ul> |
| Hung, et al., 1984                 | <ul style="list-style-type: none"> <li>• SPECT/PET/scintigraphy</li> </ul>                 | X    |      |      |      | Good    | <ul style="list-style-type: none"> <li>• Training/quality control not described</li> <li>• Precision of estimates not provided</li> </ul>  |
| Jenkins, et al., 2010              | <ul style="list-style-type: none"> <li>• Coronary CTA</li> </ul>                           | X    |      |      |      | Good    | <ul style="list-style-type: none"> <li>• Training/quality control not described</li> </ul>   |
| Johansen, et al., 2004             | <ul style="list-style-type: none"> <li>• ECG</li> <li>• SPECT/PET/scintigraphy</li> </ul>  | X    |      |      |      | Fair    | <ul style="list-style-type: none"> <li>• Source of population not described</li> <li>• Study population poorly specified</li> <li>• Version/instrumentation not specified</li> <li>• Training/quality control not described</li> <li>• Precision of estimates not provided</li> <li>• Comparability of care in international settings unclear</li> </ul> |
| Johnson, et al., 2004 <sup>d</sup> | <ul style="list-style-type: none"> <li>• CMR</li> </ul>                                    |      |      | X    |      | Fair    | <ul style="list-style-type: none"> <li>• Populations and study designs heterogeneous with varied findings</li> <li>• Comparability of care in international settings unclear</li> </ul>  |

| Study                            | NIT Modality  | KQ 1 | KQ 2 | KQ 3 | KQ 4 | Quality | Limitations to Applicability  |
|----------------------------------|---|------|------|------|------|---------|---|
| Kaminek, et al., 2001            | <ul style="list-style-type: none"> <li>SPECT/PET/scintigraphy</li> </ul>                            | X    |      |      |      | Poor    | <ul style="list-style-type: none"> <li>Source of population not described</li> <li>Study population poorly specified</li> <li>Key characteristics not reported</li> <li>Training/quality control not described</li> <li>Precision of estimates not provided</li> <li>Sequence of use of other diagnostics unclear</li> <li>Comparability of care in international settings unclear</li> </ul> |
| Kiat, et al., 1990               | <ul style="list-style-type: none"> <li>SPECT/PET/scintigraphy</li> </ul>                            | X    |      |      |      | Fair    | <ul style="list-style-type: none"> <li>Source of population not described</li> <li>Study population poorly specified</li> <li>Key characteristics not reported</li> <li>Training/quality control not described</li> <li>Precision of estimates not provided</li> <li>Data not stratified or adjusted for key predictors</li> </ul>  |
| Klem, et al., 2008               | <ul style="list-style-type: none"> <li>CMR</li> </ul>   | X    | X    |      |      | Good    | <ul style="list-style-type: none"> <li>Training/quality control not described</li> <li>Precision of estimates not provided</li> </ul>   |
| Koide, et al., 2001              | <ul style="list-style-type: none"> <li>ECG</li> </ul>   | X    |      |      |      | Good    | <ul style="list-style-type: none"> <li>Training/quality control not described</li> </ul>  |
| Langer, et al., 2009             | <ul style="list-style-type: none"> <li>CMR</li> <li>Coronary CTA</li> </ul>                         | X    |      |      |      | Good    | <ul style="list-style-type: none"> <li>Precision of estimates not provided</li> </ul>   |
| Laurienzo, et al., 1997          | <ul style="list-style-type: none"> <li>ECG</li> <li>ECHO</li> <li>SPECT/PET/scintigraphy</li> </ul> | X    |      |      | X    | Good    | <ul style="list-style-type: none"> <li>Training/quality control not described</li> <li>Precision of estimates not provided</li> <li>Data not stratified or adjusted for key predictors</li> <li>Order of testing varied across subjects and was not randomly assigned</li> </ul>  |
| Lehmkuhl, et al., 2007           | <ul style="list-style-type: none"> <li>ECG</li> <li>ECHO</li> </ul>                                 | X    |      |      | X    | Good    | <ul style="list-style-type: none"> <li>Precision of estimates not provided</li> </ul>   |
| Lewandowski, et al., 2007        | <ul style="list-style-type: none"> <li>ECG</li> <li>ECHO</li> </ul>                                 | X    |      |      |      | Good    | <ul style="list-style-type: none"> <li>Version/instrumentation not specified</li> <li>Training/quality control not described</li> <li>Comparability of care in international settings unclear</li> </ul>  |
| Lewis, et al., 1999 <sup>d</sup> | <ul style="list-style-type: none"> <li>ECHO</li> </ul>  | X    |      |      | X    | Good    | <ul style="list-style-type: none"> <li>Precision of estimates not provided</li> </ul>   |
| Lewis, et al., 2005 <sup>d</sup> | <ul style="list-style-type: none"> <li>ECG</li> </ul>   | X    | X    | X    |      | Good    | <ul style="list-style-type: none"> <li>Key characteristics not reported</li> <li>Time from results to treatment not reported</li> </ul>   |
| Lu, et al., 2010                 | <ul style="list-style-type: none"> <li>ECG</li> <li>ECHO</li> <li>SPECT/PET/scintigraphy</li> </ul> | X    | X    |      | X    | Good    | <ul style="list-style-type: none"> <li>Training/quality control not described</li> <li>Precision of estimates not provided</li> </ul>   |



| Study                   | NIT Modality   | KQ 1 | KQ 2 | KQ 3 | KQ 4 | Quality | Limitations to Applicability   |
|-------------------------|--|------|------|------|------|---------|--|
| Maffei, et al., 2010    | <ul style="list-style-type: none"> <li>ECG</li> </ul>                                  | X    |      |      |      | Fair    | <ul style="list-style-type: none"> <li>Source of population not described</li> <li>Precision estimates not provided</li> <li>Sequence of use of other diagnostics unclear</li> <li>Comparability of care in international settings unclear</li> </ul>  |
| Majstorov, et al., 2005 | <ul style="list-style-type: none"> <li>SPECT/PET/scintigraphy</li> </ul>               | X    |      |      |      | Poor    | <ul style="list-style-type: none"> <li>Source of population not described</li> <li>Study population poorly specified</li> <li>Key characteristics not reported</li> <li>Training/quality control not described</li> <li>Precision of estimates not provided</li> </ul>   |
| Mak, et al., 1995       | <ul style="list-style-type: none"> <li>SPECT/PET/scintigraphy</li> </ul>               | X    |      |      |      | Good    | <ul style="list-style-type: none"> <li>Study population poorly specified</li> <li>Precision of estimates not provided</li> </ul>   |
| Marwick, et al., 1993   | <ul style="list-style-type: none"> <li>ECHO</li> <li>SPECT/PET/scintigraphy</li> </ul> | X    |      |      |      | Fair    | <ul style="list-style-type: none"> <li>Study population poorly specified</li> <li>Key characteristics not reported</li> <li>Training/quality control not described</li> <li>Precision of estimates not provided</li> <li>Data not stratified or adjusted for key predictors</li> <li>Order of testing varied across subjects and was not randomly assigned</li> </ul>  |
| Marwick, et al., 1995   | <ul style="list-style-type: none"> <li>ECG</li> <li>ECHO</li> </ul>                    | X    | X    |      |      | Fair    | <ul style="list-style-type: none"> <li>Key characteristics not reported</li> <li>Version/instrumentation not specified</li> <li>Training/quality control not described</li> <li>Comparability of care in international settings unclear</li> </ul>   |
| Masini, et al., 1988    | <ul style="list-style-type: none"> <li>ECG</li> <li>ECHO</li> </ul>                    | X    |      |      |      | Fair    | <ul style="list-style-type: none"> <li>Source of population not described</li> <li>Key characteristics not reported</li> <li>Version/instrumentation not specified</li> <li>Training/quality control not described</li> <li>Precision of estimates not provided</li> <li>Sequence of use of other diagnostics unclear</li> <li>Order of testing varied across subjects and was not randomly assigned</li> <li>Comparability of care in international settings unclear</li> </ul> |
| Mazeika, et al., 1992   | <ul style="list-style-type: none"> <li>ECHO</li> </ul>                                 | X    |      |      |      | Fair    | <ul style="list-style-type: none"> <li>Study population poorly specified</li> <li>Key characteristics not reported</li> <li>Precision of estimates not provided</li> <li>Data not stratified or adjusted for key predictors</li> </ul>   |
| Merkle, et al., 2010    | <ul style="list-style-type: none"> <li>CMR</li> </ul>                                  | X    |      |      |      | Good    | <ul style="list-style-type: none"> <li>Training/quality control not described</li> <li>Data not stratified or adjusted for key predictors</li> </ul>   |

| Study                             | NIT Modality  | KQ 1 | KQ 2 | KQ 3 | KQ 4 | Quality | Limitations to Applicability   |
|-----------------------------------|---|------|------|------|------|---------|--|
| Michaelides, et al., 2007         | <ul style="list-style-type: none"> <li>ECG</li> </ul>                                 | X    |      |      |      | Fair    | <ul style="list-style-type: none"> <li>Source of population not described</li> <li>Study population poorly specified</li> <li>Version/instrumentation not specified</li> <li>Training/quality control not described</li> <li>Precision of estimates not provided</li> <li>Comparability of care in international settings unclear</li> </ul>   |
| Mieres, et al., 2007              | <ul style="list-style-type: none"> <li>ECG</li> <li>SPECT/PET/scintigraphy</li> </ul> | X    |      | X    |      | Fair    | <ul style="list-style-type: none"> <li>Source of population not described</li> <li>Version/instrumentation not specified</li> <li>Training/quality control not described</li> <li>Failure to test “normals” or subset with gold standard</li> <li>Precision of estimates not provided</li> <li>Time from results to treatment not reported</li> <li>Comparability of care in international settings unclear</li> </ul> |
| Miller, et al., 2001              | <ul style="list-style-type: none"> <li>ECG</li> <li>SPECT/PET/scintigraphy</li> </ul> | X    |      |      |      | Fair    | <ul style="list-style-type: none"> <li>Training/quality control not described</li> <li>Precision of estimates not provided</li> <li>Data not stratified or adjusted for key predictors</li> </ul>  |
| Mohiuddin, et al., 1996           | <ul style="list-style-type: none"> <li>SPECT/PET/scintigraphy</li> </ul>              | X    |      |      | X    | Good    | <ul style="list-style-type: none"> <li>Training/quality control not described</li> </ul>   |
| Morise, et al., 2000              | <ul style="list-style-type: none"> <li>ECG</li> </ul>                                 | X    |      |      |      | Good    | <ul style="list-style-type: none"> <li>None</li> </ul>   |
| Morise, et al., 1995-A            | <ul style="list-style-type: none"> <li>ECG</li> </ul>                                 | X    |      |      |      | Fair    | <ul style="list-style-type: none"> <li>Training/quality control not described</li> <li>Precision of estimates not provided</li> </ul>  |
| Morise, et al., 1995-B            | <ul style="list-style-type: none"> <li>SPECT/PET/scintigraphy</li> </ul>              | X    |      |      |      | Poor    | <ul style="list-style-type: none"> <li>Study population poorly specified</li> <li>Training/quality control not described</li> <li>Precision of estimates not provided</li> <li>Data not stratified or adjusted for key predictors</li> </ul>   |
| Morise, et al., 2002              | <ul style="list-style-type: none"> <li>ECG</li> </ul>                                 |      |      | X    |      | Good    | <ul style="list-style-type: none"> <li>Precision of estimates not provided</li> <li>Comparability of care in international settings unclear</li> </ul>   |
| Morise, et al., 2004 <sup>d</sup> | <ul style="list-style-type: none"> <li>ECG</li> </ul>                                 |      |      | X    |      | Fair    | <ul style="list-style-type: none"> <li>Source of population not described</li> <li>Study population poorly specified</li> <li>Version/instrumentation not specified</li> <li>Training/quality control not described</li> <li>Precision of estimates not provided</li> <li>Sequence of use of other diagnostics unclear</li> <li>Comparability of care in international settings unclear</li> </ul>                     |

| Study                    | NIT Modality  | KQ 1 | KQ 2 | KQ 3 | KQ 4 | Quality | Limitations to Applicability   |
|--------------------------|---|------|------|------|------|---------|--|
| Nascimento, et al., 2008 | • ECG   | X    |      |      |      | Fair    | <ul style="list-style-type: none"> <li>• Training/quality control not described</li> <li>• Data not stratified or adjusted for key predictors</li> <li>• Sequence of use of other diagnostics unclear</li> </ul>   |
| Ozdemir, et al., 2002    | • ECG   | X    |      |      |      | Good    | <ul style="list-style-type: none"> <li>• Training/quality control not described</li> <li>• Precision of estimates not provided</li> <li>• Data not stratified or adjusted for key predictors</li> </ul>  |
| Pundziute, et al., 2008  | • Coronary CTA  | X    |      |      |      | Fair    | <ul style="list-style-type: none"> <li>• Source of population not described</li> <li>• Sequence of use of other diagnostics unclear</li> <li>• Comparability of care in international settings unclear</li> </ul>  |
| Raman, et al., 2008      | <ul style="list-style-type: none"> <li>• ECG</li> <li>• SPECT/PET/scintigraphy</li> <li>• CMR</li> </ul>  | X    |      | X    |      | Fair    | <ul style="list-style-type: none"> <li>• Source of population not described</li> <li>• Training/quality control not described</li> <li>• Comparability of care in international settings unclear</li> </ul>  |
| Richardson, et al., 1995 | • ECG   | X    |      |      |      | Good    | <ul style="list-style-type: none"> <li>• Training/quality control not described</li> <li>• Precision of estimates not provided</li> <li>• Data not stratified or adjusted for key predictors</li> </ul>  |
| Robert, et al., 1991     | • ECG   | X    |      |      |      | Fair    | <ul style="list-style-type: none"> <li>• Source of population not described</li> <li>• Key characteristics not reported</li> <li>• Training/quality control not described</li> <li>• Precision of estimates not provided</li> <li>• Comparability of care in international settings unclear</li> </ul> |
| Roger, et al., 1997      | • ECHO  | X    |      |      |      | Poor    | <ul style="list-style-type: none"> <li>• Source of population not described</li> <li>• Study population poorly specified</li> <li>• Training/quality control not described</li> </ul>  |
| Rollan, et al., 2002     | <ul style="list-style-type: none"> <li>• ECG</li> <li>• ECHO</li> <li>• SPECT/PET/scintigraphy</li> </ul> | X    |      |      |      | Fair    | <ul style="list-style-type: none"> <li>• Source of population not described</li> <li>• Training/quality control not described</li> <li>• Sequence of use of other diagnostics unclear</li> <li>• Comparability of care in international settings unclear</li> </ul>                                    |
| San Roman, et al., 1998  | <ul style="list-style-type: none"> <li>• ECG</li> <li>• ECHO</li> <li>• SPECT/PET/scintigraphy</li> </ul> | X    |      |      |      | Good    | <ul style="list-style-type: none"> <li>• Source of population not described</li> <li>• Study population poorly specified</li> <li>• Version/instrumentation not specified</li> <li>• Precision of estimates not provided</li> <li>• Comparability of care in international settings unclear</li> </ul> |

| Study                           | NIT Modality  | KQ 1 | KQ 2 | KQ 3 | KQ 4 | Quality | Limitations to Applicability   |
|---------------------------------|---|------|------|------|------|---------|--|
| Sanfilippo, et al., 2005        | <ul style="list-style-type: none"> <li>• ECG</li> <li>• ECHO</li> </ul>                   |      |      | X    |      | Fair    | <ul style="list-style-type: none"> <li>• Version/instrumentation not specified</li> <li>• Training/quality control not described</li> <li>• Screening and diagnostic uses mixed</li> <li>• Gold standard not applied</li> <li>• Correlational data only</li> <li>• Failure to test “normals” or subset with gold standard</li> <li>• Precision of estimates not provided</li> <li>• Data not stratified or adjusted for key predictors</li> <li>• Sequence of use of other diagnostics unclear</li> <li>• Comparability of care in international settings unclear</li> </ul> |
| Santana-Boado, et al., 1998     | <ul style="list-style-type: none"> <li>• ECG</li> <li>• SPECT/PET/scintigraphy</li> </ul> | X    |      |      |      | Fair    | <ul style="list-style-type: none"> <li>• Source of population not described</li> <li>• Study population poorly specified</li> <li>• Comparability of care in international settings unclear</li> </ul>   |
| Sawada, et al., 1989            | <ul style="list-style-type: none"> <li>• ECHO</li> </ul>                                  | X    |      |      |      | Good    | <ul style="list-style-type: none"> <li>• Source of population not described</li> <li>• Precision of estimates not provided</li> <li>• Comparability of care in international settings unclear</li> </ul>   |
| Schupbach, et al., 2008         | <ul style="list-style-type: none"> <li>• ECG</li> </ul>                                   | X    |      |      |      | Fair    | <ul style="list-style-type: none"> <li>• Source of population not described</li> <li>• Study population poorly specified</li> <li>• Training/quality control not described</li> <li>• Precision of estimates not provided</li> <li>• Comparability of care in international settings unclear</li> </ul>  |
| Severi, et al., 1994            | <ul style="list-style-type: none"> <li>• ECG</li> <li>• ECHO</li> </ul>                   | X    |      |      |      | Good    | <ul style="list-style-type: none"> <li>• Precision of estimates not provided</li> <li>• Data not stratified or adjusted for key predictors</li> <li>• Order of testing varied across subjects and was not randomly assigned</li> </ul>   |
| Sharir, et al., 2000            | <ul style="list-style-type: none"> <li>• SPECT/PET/scintigraphy</li> </ul>                | X    |      |      |      | Poor    | <ul style="list-style-type: none"> <li>• Source of population not described</li> <li>• Study population poorly specified</li> <li>• Key characteristics not reported</li> <li>• Training/quality control not described</li> <li>• Precision of estimates not provided</li> <li>• Data not stratified or adjusted for key predictors</li> </ul>   |
| Shaw, et al., 2011 <sup>e</sup> | <ul style="list-style-type: none"> <li>• ECG</li> <li>• SPECT/PET/scintigraphy</li> </ul> |      |      | X    |      | Fair    | <ul style="list-style-type: none"> <li>• Gold standard not applied</li> </ul>  |

| Study                   | NIT Modality   | KQ 1 | KQ 2 | KQ 3 | KQ 4 | Quality | Limitations to Applicability  |
|-------------------------|--|------|------|------|------|---------|---|
| Shi, et al., 2007       | <ul style="list-style-type: none"> <li>SPECT/PET/scintigraphy</li> </ul>               | X    |      |      |      | Poor    | <ul style="list-style-type: none"> <li>Source of population not described</li> <li>Study population poorly specified</li> <li>Key characteristics not reported</li> <li>Training/quality control not described</li> <li>Failure to test “normals” or subset with gold standard</li> <li>Precision of estimates not provided</li> <li>Data not stratified or adjusted for key predictors</li> <li>Comparability of care in international settings unclear</li> </ul> |
| Shin, et al., 2003      | <ul style="list-style-type: none"> <li>ECHO</li> </ul>                                 | X    | X    |      |      | Fair    | <ul style="list-style-type: none"> <li>Source of population not described</li> <li>Training/quality control not described</li> <li>Precision of estimates not provided</li> <li>Comparability of care in international settings unclear</li> </ul>  |
| Shivalkar, et al., 2007 | <ul style="list-style-type: none"> <li>Coronary CTA</li> </ul>                         | X    |      |      |      | Fair    | <ul style="list-style-type: none"> <li>Source of population not described</li> <li>Study population poorly specified</li> <li>Training/quality control not described</li> <li>Comparability of care in international settings unclear</li> </ul>  |
| Siegler, et al., 2011   | <ul style="list-style-type: none"> <li>ECG</li> <li>SPECT/PET/scintigraphy</li> </ul>  | X    | X    |      |      | Fair    | <ul style="list-style-type: none"> <li>Gold standard not applied</li> <li>Precision of estimates not provided</li> </ul>  |
| Sinha, et al., 2008     | <ul style="list-style-type: none"> <li>ECG</li> </ul>                                  | X    |      |      |      | Poor    | <ul style="list-style-type: none"> <li>Key characteristics not reported</li> <li>Training/quality control not described</li> <li>Precision of estimates not provided</li> <li>Data not stratified or adjusted for key predictors</li> </ul>   |
| Sketch, et al., 1975    | <ul style="list-style-type: none"> <li>ECG</li> </ul>                                  | X    |      |      |      | Poor    | <ul style="list-style-type: none"> <li>Source of population not described</li> <li>Study population poorly specified</li> <li>Precision of estimates not provided</li> <li>Sequence of use of other diagnostics unclear</li> <li>Comparability of care in international settings unclear</li> </ul>   |
| Slavich, et al., 1996   | <ul style="list-style-type: none"> <li>ECHO</li> <li>SPECT/PET/scintigraphy</li> </ul> | X    |      |      |      | Fair    | <ul style="list-style-type: none"> <li>Source of population not described</li> <li>Study population poorly specified</li> <li>Key characteristics not reported</li> <li>Training/quality control not described</li> <li>Precision of estimates not provided</li> <li>Comparability of care in international settings unclear</li> </ul>   |

| Study                   | NIT Modality   | KQ 1 | KQ 2 | KQ 3 | KQ 4 | Quality | Limitations to Applicability  |
|-------------------------|--|------|------|------|------|---------|---|
| Slomka, et al., 2007    | <ul style="list-style-type: none"> <li>SPECT/PET/scintigraphy</li> </ul>               | X    |      |      |      | Poor    | <ul style="list-style-type: none"> <li>Source of population not described</li> <li>Study population poorly specified</li> <li>Training/quality control not described</li> <li>Precision of estimates not provided</li> <li>Comparability of care in international settings unclear</li> </ul>   |
| Slomka, et al., 2005    | <ul style="list-style-type: none"> <li>SPECT/PET/scintigraphy</li> </ul>               | x    |      |      |      | Fair    | <ul style="list-style-type: none"> <li>Source of population not described</li> <li>Study population poorly specified</li> <li>Training/quality control not described</li> <li>Precision of estimates not provided</li> <li>Data not stratified or adjusted for key predictors</li> <li>Comparability of care in international settings unclear</li> </ul>   |
| Svart, et al., 2010     | <ul style="list-style-type: none"> <li>ECG</li> </ul>                                  | X    |      |      |      | Fair    | <ul style="list-style-type: none"> <li>Study population poorly specified</li> <li>Key characteristics not reported</li> <li>Training/quality control not described</li> <li>Precision of estimates not provided</li> </ul>  |
| Taillefer, et al., 1997 | <ul style="list-style-type: none"> <li>SPECT/PET/scintigraphy</li> </ul>               | X    |      |      |      | Good    | <ul style="list-style-type: none"> <li>Study population poorly specified</li> <li>Training/quality control not described</li> <li>Precision of estimates not provided</li> </ul>  |
| Takeuchi, et al., 1996  | <ul style="list-style-type: none"> <li>ECHO</li> <li>SPECT/PET/scintigraphy</li> </ul> | X    |      |      |      | Fair    | <ul style="list-style-type: none"> <li>Source of population not described</li> <li>Study population poorly specified</li> <li>Key characteristics not reported</li> <li>Training/quality control not described</li> <li>Precision of estimates not provided</li> <li>Comparability of care in international settings unclear</li> </ul>   |
| Travin, et al., 2000    | <ul style="list-style-type: none"> <li>SPECT/PET/scintigraphy</li> </ul>               | X    |      |      |      | Fair    | <ul style="list-style-type: none"> <li>Source of population not described</li> <li>Study population poorly specified</li> <li>Precision of estimates not provided</li> <li>Comparability of care in international settings unclear</li> </ul>   |
| Van Train, et al., 1994 | <ul style="list-style-type: none"> <li>SPECT/PET/scintigraphy</li> </ul>               | X    |      |      |      | Poor    | <ul style="list-style-type: none"> <li>Source of population not described</li> <li>Study population poorly specified</li> <li>Key characteristics not reported</li> <li>Training/quality control not described</li> <li>Precision of estimates not provided</li> <li>Data not stratified or adjusted for key predictors</li> <li>Order of testing varied across subjects and was not randomly assigned</li> </ul> |

| Study                  | NIT Modality                 | KQ 1 | KQ 2 | KQ 3 | KQ 4 | Quality | Limitations to Applicability   |
|------------------------|------------------------------|------|------|------|------|---------|--|
| Vashist, et al., 2007  | • SPECT/PET/<br>scintigraphy | X    | X    | X    |      | Fair    | <ul style="list-style-type: none"> <li>• Source of population not described</li> <li>• Training/quality control not described</li> <li>• Comparability of care in international settings unclear</li> </ul>  |
| Weiner, et al., 1985   | • ECG                        | X    |      |      |      | Poor    | <ul style="list-style-type: none"> <li>• Source of population not described</li> <li>• Study population poorly specified</li> <li>• Key characteristics not reported</li> <li>• Training/quality control not described</li> <li>• Precision of estimates not provided</li> <li>• Data not stratified or adjusted for key predictors</li> </ul>                 |
| Weiner, et al., 1979   | • ECG                        | X    |      |      |      | Good    | <ul style="list-style-type: none"> <li>• Study population poorly specified</li> <li>• Key characteristics not reported</li> <li>• Training/quality control not described</li> <li>• Precision of estimates not provided</li> </ul>   |
| Weustink, et al., 2007 | • Coronary CTA               |      |      |      | X    | Fair    | <ul style="list-style-type: none"> <li>• Source of population not described</li> <li>• Precision of estimates not provided</li> <li>• Comparability of care in international settings unclear</li> </ul>   |
| Weustink, et al., 2009 | • Coronary CTA               |      |      |      | X    | Good    | <ul style="list-style-type: none"> <li>• None</li> </ul>   |
| Williams, et al., 1994 | • ECHO                       | X    |      |      |      | Fair    | <ul style="list-style-type: none"> <li>• Key characteristics not reported</li> <li>• Training/quality control not described</li> <li>• Precision of estimates not provided</li> <li>• Data not stratified or adjusted for key predictors</li> </ul>  |
| Wolak, et al., 2008    | • SPECT/PET/<br>scintigraphy | X    |      |      |      | Fair    | <ul style="list-style-type: none"> <li>• Source of population not described</li> <li>• Comparability of care in international settings unclear</li> </ul>  |
| Wong, et al., 2001     | • ECG                        |      |      | X    |      | Poor    | <ul style="list-style-type: none"> <li>• Source of population not described</li> <li>• Study population poorly specified</li> <li>• Key characteristics not reported</li> <li>• Training/quality control not described</li> <li>• Precision of estimates not provided</li> <li>• Time from results to treatment not reported</li> </ul>                        |
| Yamauchi, et al., 1985 | • ECG                        | X    |      |      |      | Fair    | <ul style="list-style-type: none"> <li>• Source of population not described</li> <li>• Key characteristics not reported</li> <li>• Training/quality control not described</li> <li>• Precision of estimates not provided</li> <li>• Sequence of use of other diagnostics unclear</li> <li>• Comparability of care in international settings unclear</li> </ul> |

| Study              | NIT Modality  | KQ 1 | KQ 2 | KQ 3 | KQ 4 | Quality | Limitations to Applicability  |
|--------------------|---|------|------|------|------|---------|---|
| Yeih, et al., 2007 | <ul style="list-style-type: none"> <li>• ECG</li> <li>• SPECT/PET/scintigraphy</li> </ul> | X    | X    |      |      | Fair    | <ul style="list-style-type: none"> <li>• Source of population not described</li> <li>• Study population poorly specified</li> <li>• Precision of estimates not provided</li> <li>• Comparability of care in international settings unclear</li> </ul> |

<sup>a</sup>Related methods article: Dewey, et al., 2006 (refer to Appendix C for full citation).

<sup>b</sup>Related methods article: Miller, et al., 2009 (refer to Appendix C for full citation).

<sup>c</sup>Related methods articles: Meijboom, et al., 2007 and Meijboom, et al., 2008 (refer to Appendix C for full citations).

<sup>d</sup>Related methods article: Merz, et al., 1999 (refer to Appendix C for full citation).

<sup>e</sup>Related methods article: Mieres, et al., 2009 (refer to Appendix C for full citation).

Abbreviations: CAD = coronary artery disease; CMR =cardiac magnetic resonance; CTA = computed tomography angiography; ECG = exercise/stress electrocardiogram; ECHO = echocardiogram; KQ = Key Question; PET = positron emission tomography; SPECT = single proton emission computed tomography

**Table D-2. QUADAS tool for quality assessment of diagnostic accuracy<sup>a</sup>**

| Study                   | Patient Representation | Selection Criteria | Reference Standard | Time Interval | Sample for Verification | Reference test uniformly applied | Reference test independently performed | Index Test Replicability | Reference Test Replicability | Index Test Blinded Interpretation | Reference Test Blinded Interpretation | Clinical Data Availability | Uninterpretable or Intermediate Test Reporting | Study Withdrawal Description |
|-------------------------|------------------------|--------------------|--------------------|---------------|-------------------------|----------------------------------|--|--------------------------|------------------------------|-----------------------------------|---------------------------------------|----------------------------|--|------------------------------|
| Abramson, et al., 2000  | N                      | Y                  | Y                  | Y             | Y                       | Y                                | Y                                      | Y                        | Y                            | Y                                 | Y                                     | Y                          | N  | N                            |
| Agati, et al., 1992     | U                      | N                  | Y                  | Y             | Y                       | Y                                | Y                                      | Y                        | Y                            | Y                                 | Y                                     | U                          | N  | N                            |
| Barolsky, et al., 1979  | U                      | N                  | Y                  | N             | Y                       | Y                                | Y                                      | Y                        | Y                            | U                                 | U                                     | U                          | N  | N                            |
| Bjornstad, et al., 1995 | N                      | Y                  | Y                  | Y             | Y                       | Y                                | Y                                      | Y                        | Y                            | Y                                 | Y                                     | Y                          | N  | Y                            |
| Bokhari, et al., 2008   | N                      | Y                  | Y                  | Y             | N                       | N                                | N                                      | Y                        | Y                            | N                                 | N                                     | Y                          | N  | N                            |



| Study                                | Patient Representation | Selection Criteria | Reference Standard | Time Interval | Sample for Verification | Reference test uniformly applied | Reference test independently performed | Index Test Replicability | Reference Test Replicability | Index Test Blinded Interpretation | Reference Test Blinded Interpretation | Clinical Data Availability | Uninterpretable or Intermediate Test Reporting | Study Withdrawal Description |
|--------------------------------------|------------------------|--------------------|--------------------|---------------|-------------------------|----------------------------------|--|--------------------------|------------------------------|-----------------------------------|---------------------------------------|----------------------------|--|------------------------------|
| Burgi Wegmann, et al., 2003          | U                      | Y                  | Y                  | U             | Y                       | Y                                | Y                                      | N                        | U                            | U                                 | U                                     | U                          | N  | N                            |
| Chae, et al., 1993                   | Y                      | Y                  | Y                  | Y             | Y                       | Y                                | Y                                      | Y                        | Y                            | Y                                 | Y                                     | Y                          | Y  | N                            |
| Cin, et al., 2000                    | Y                      | Y                  | Y                  | Y             | N                       | Y                                | Y                                      | Y                        | Y                            | U                                 | U                                     | Y                          | N  | Y                            |
| Coelho-Filho, et al., 2011           | Y                      | N                  | Y                  | N             | N                       | N                                | Y                                      | Y                        | Y                            | Y                                 | Y                                     | U                          | Y  | Y                            |
| DePasquale, et al., 1988             | U                      | N                  | Y                  | Y             | Y                       | Y                                | Y                                      | Y                        | Y                            | Y                                 | U                                     | U                          | N  | N                            |
| Dewey, et al., 2008 <sup>b</sup>     | N                      | Y                  | Y                  | Y             | Y                       | Y                                | Y                                      | Y                        | Y                            | Y                                 | U                                     | N                          | U  | N                            |
| Dewey, et al., 2010 <sup>c</sup>     | Y                      | Y                  | Y                  | Y             | Y                       | Y                                | Y                                      | Y                        | Y                            | Y                                 | Y                                     | N                          | Y  | N                            |
| Dharampal, et al., 2011 <sup>d</sup> | Y                      | Y                  | Y                  | Y             | Y                       | Y                                | Y                                      | Y                        | N                            | U                                 | U                                     | U                          | N  | Y                            |
| Dionisopoulos, et al., 1997          | U                      | N                  | Y                  | Y             | Y                       | Y                                | Y                                      | Y                        | Y                            | Y                                 | Y                                     | Y                          | N  | N                            |
| Dodi, et al., 2001                   | Y                      | Y                  | N                  | Y             | Y                       | Y                                | Y                                      | Y                        | Y                            | U                                 | U                                     | Y                          | N  | N                            |
| Doyle, et al., 2003 <sup>e</sup>     | Y                      | Y                  | Y                  | U             | Y                       | Y                                | Y                                      | Y                        | Y                            | Y                                 | Y                                     | U                          | N  | Y                            |
| Doyle, et al., 2010 <sup>e</sup>     | U                      | Y                  | Y                  | Y             | Y                       | Y                                | Y                                      | Y                        | N                            | N                                 | Y                                     | U                          | N  | N                            |

| Study                  | Patient Representation | Selection Criteria | Reference Standard | Time Interval | Sample for Verification | Reference test uniformly applied | Reference test independently performed | Index Test Replicability | Reference Test Replicability | Index Test Blinded Interpretation | Reference Test Blinded Interpretation | Clinical Data Availability | Uninterpretable or Intermediate Test Reporting | Study Withdrawal Description |
|------------------------|------------------------|--------------------|--------------------|---------------|-------------------------|----------------------------------|--|--------------------------|------------------------------|-----------------------------------|---------------------------------------|----------------------------|--|------------------------------|
| Elhendy, et al., 1997  | U                      | Y                  | Y                  | Y             | Y                       | Y                                | Y                                      | Y                        | U                            | Y                                 | U                                     | N                          | N  | N                            |
| Elhendy, et al., 2006  | U                      | Y                  | Y                  | Y             | Y                       | Y                                | U                                      | Y                        | U                            | Y                                 | N                                     | U                          | N  | N                            |
| Elhendy, et al., 1998  | Y                      | N                  | Y                  | Y             | Y                       | Y                                | U                                      | Y                        | Y                            | Y                                 | Y                                     | Y                          | N  | N                            |
| Emmett, et al., 2002   | U                      | Y                  | Y                  | U             | Y                       | Y                                | Y                                      | Y                        | U                            | Y                                 | Y                                     | N                          | U  | N                            |
| Faisal, et al., 2007   | U                      | N                  | Y                  | U             | N                       | N                                | N                                      | Y                        | U                            | U                                 | N                                     | U                          | N  | N                            |
| Friedman, et al., 1982 | U                      | N                  | Y                  | U             | Y                       | Y                                | Y                                      | Y                        | U                            | Y                                 | Y                                     | U                          | Y  | N                            |
| Gebker, et al., 2010   | U                      | Y                  | Y                  | Y             | Y                       | Y                                | Y                                      | Y                        | Y                            | Y                                 | Y                                     | U                          | Y  | Y                            |
| Gentile, et al., 2001  | Y                      | U                  | Y                  | Y             | Y                       | Y                                | Y                                      | Y                        | Y                            | Y                                 | Y                                     | Y                          | N  | N                            |
| Guiteras, et al., 1982 | U                      | Y                  | Y                  | Y             | Y                       | Y                                | U                                      | Y                        | U                            | U                                 | U                                     | U                          | Y  | N                            |
| Gulati, et al., 2004   | U                      | Y                  | Y                  | U             | Y                       | Y                                | Y                                      | Y                        | U                            | Y                                 | U                                     | U                          | N  | N                            |
| Hlatky, et al., 1984   | U                      | Y                  | Y                  | U             | Y                       | Y                                | U                                      | Y                        | U                            | U                                 | U                                     | U                          | Y  | N                            |
| Ho, et al., 1998       | U                      | Y                  | Y                  | Y             | Y                       | Y                                | Y                                      | Y                        | Y                            | Y                                 | U                                     | U                          | N  | Y                            |

| Study                              | Patient Representation | Selection Criteria | Reference Standard | Time Interval | Sample for Verification | Reference test uniformly applied | Reference test independently performed | Index Test Replicability | Reference Test Replicability | Index Test Blinded Interpretation | Reference Test Blinded Interpretation | Clinical Data Availability | Uninterpretable or Intermediate Test Reporting | Study Withdrawal Description |
|------------------------------------|------------------------|--------------------|--------------------|---------------|-------------------------|----------------------------------|--|--------------------------|------------------------------|-----------------------------------|---------------------------------------|----------------------------|--|------------------------------|
| Hoilund-Carlsen, et al., 2005      | U                      | N                  | Y                  | Y             | Y                       | Y                                | U                                      | Y                        | U                            | Y                                 | U                                     | U                          | N  | N                            |
| Hoilund-Carlsen, et al., 2007      | Y                      | Y                  | Y                  | Y             | Y                       | Y                                | Y                                      | Y                        | N                            | Y                                 | Y                                     | U                          | Y  | Y                            |
| Hosokawa, et al., 2008             | N                      | N                  | Y                  | U             | Y                       | Y                                | Y                                      | Y                        | U                            | Y                                 | Y                                     | N                          | Y  | Y                            |
| Hung, et al., 1984                 | Y                      | Y                  | Y                  | Y             | Y                       | Y                                | Y                                      | Y                        | Y                            | Y                                 | Y                                     | U                          | Y  | N                            |
| Jenkins, et al., 2010              | Y                      | Y                  | Y                  | Y             | Y                       | Y                                | Y                                      | Y                        | Y                            | Y                                 | Y                                     | U                          | Y  | Y                            |
| Johansen, et al., 2004             | U                      | Y                  | Y                  | U             | N                       | N                                | U                                      | Y                        | N                            | Y                                 | U                                     | N                          | N  | N                            |
| Johnson, et al., 2004 <sup>e</sup> | N                      | Y                  | Y                  | U             | Y                       | Y                                | Y                                      | Y                        | Y                            | N                                 | Y                                     | U                          | N  | N                            |
| Kaminek, et al., 2001              | U                      | N                  | Y                  | U             | Y                       | Y                                | Y                                      | U                        | N                            | U                                 | U                                     | U                          | N  | N                            |
| Kiat, et al., 1990                 | U                      | N                  | Y                  | Y             | N                       | Y                                | Y                                      | Y                        | Y                            | Y                                 | Y                                     | U                          | N  | N                            |
| Klem, et al., 2008                 | Y                      | Y                  | Y                  | Y             | Y                       | Y                                | Y                                      | Y                        | Y                            | Y                                 | Y                                     | Y                          | N  | Y                            |
| Koide, et al., 2001                | Y                      | Y                  | Y                  | Y             | Y                       | Y                                | Y                                      | Y                        | Y                            | Y                                 | U                                     | Y                          | N  | Y                            |
| Langer, et al., 2009               | U                      | Y                  | Y                  | Y             | Y                       | Y                                | Y                                      | N                        | Y                            | Y                                 | Y                                     | U                          | Y  | Y                            |

| Study                            | Patient Representation | Selection Criteria | Reference Standard | Time Interval | Sample for Verification | Reference test uniformly applied | Reference test independently performed | Index Test Replicability | Reference Test Replicability | Index Test Blinded Interpretation | Reference Test Blinded Interpretation | Clinical Data Availability | Uninterpretable or Intermediate Test Reporting | Study Withdrawal Description |
|----------------------------------|------------------------|--------------------|--------------------|---------------|-------------------------|----------------------------------|--|--------------------------|------------------------------|-----------------------------------|---------------------------------------|----------------------------|--|------------------------------|
| Laurienzo, et al., 1997          | Y                      | Y                  | Y                  | Y             | Y                       | Y                                | Y                                      | Y                        | Y                            | U                                 | Y                                     | U                          | N  | Y                            |
| Lehmkuhl, et al., 2007           | Y                      | Y                  | Y                  | Y             | Y                       | Y                                | Y                                      | Y                        | Y                            | Y                                 | Y                                     | Y                          | Y  | Y                            |
| Lewandowski, et al., 2007        | Y                      | Y                  | Y                  | Y             | Y                       | Y                                | Y                                      | Y                        | U                            | Y                                 | U                                     | U                          | N  | N                            |
| Lewis, et al., 1999 <sup>e</sup> | Y                      | Y                  | Y                  | U             | Y                       | Y                                | Y                                      | Y                        | Y                            | Y                                 | Y                                     | Y                          | Y  | N                            |
| Lewis, et al., 2005 <sup>e</sup> | Y                      | Y                  | Y                  | U             | Y                       | Y                                | U                                      | Y                        | Y                            | Y                                 | Y                                     | Y                          | Y  | Y                            |
| Lu, et al., 2010                 | Y                      | Y                  | Y                  | Y             | Y                       | Y                                | Y                                      | Y                        | Y                            | U                                 | U                                     | U                          | N  | Y                            |
| Maffei, et al., 2010             | Y                      | Y                  | Y                  | U             | Y                       | Y                                | U                                      | Y                        | Y                            | Y                                 | U                                     | U                          | Y  | Y                            |
| Majstorov, et al., 2005          | U                      | N                  | Y                  | Y             | Y                       | Y                                | U                                      | U                        | N                            | Y                                 | Y                                     | U                          | N  | N                            |
| Mak, et al., 1995                | Y                      | N                  | Y                  | Y             | Y                       | Y                                | Y                                      | Y                        | Y                            | Y                                 | Y                                     | Y                          | Y  | U                            |
| Marwick, et al., 1993            | U                      | N                  | Y                  | Y             | Y                       | Y                                | Y                                      | Y                        | Y                            | Y                                 | Y                                     | U                          | N  | N                            |
| Marwick, et al., 1995            | U                      | Y                  | Y                  | N             | Y                       | Y                                | Y                                      | Y                        | U                            | Y                                 | Y                                     | U                          | Y  | N                            |
| Masini, et al., 1988             | U                      | U                  | Y                  | U             | Y                       | Y                                | Y                                      | U                        | U                            | U                                 | U                                     | U                          | Y  | N                            |

| Study                             | Patient Representation | Selection Criteria | Reference Standard | Time Interval | Sample for Verification | Reference test uniformly applied | Reference test independently performed | Index Test Replicability | Reference Test Replicability | Index Test Blinded Interpretation | Reference Test Blinded Interpretation | Clinical Data Availability | Uninterpretable or Intermediate Test Reporting | Study Withdrawal Description |
|-----------------------------------|------------------------|--------------------|--------------------|---------------|-------------------------|----------------------------------|--|--------------------------|------------------------------|-----------------------------------|---------------------------------------|----------------------------|--|------------------------------|
| Mazeika, et al., 1992             | Y                      | N                  | Y                  | U             | Y                       | Y                                | Y                                      | Y                        | Y                            | Y                                 | Y                                     | U                          | N  | N                            |
| Merkle, et al., 2010              | Y                      | Y                  | Y                  | U             | Y                       | Y                                | Y                                      | Y                        | Y                            | Y                                 | Y                                     | U                          | N  | N                            |
| Michaelides, et al., 2007         | U                      | N                  | Y                  | Y             | Y                       | Y                                | Y                                      | Y                        | U                            | Y                                 | Y                                     | U                          | N  | N                            |
| Mieres, et al., 2007              | U                      | Y                  | Y                  | Y             | N                       | Y                                | Y                                      | N                        | N                            | Y                                 | Y                                     | N                          | N  | Y                            |
| Miller, et al., 2001              | Y                      | Y                  | Y                  | Y             | N                       | Y                                | Y                                      | Y                        | Y                            | U                                 | U                                     | Y                          | N  | N                            |
| Mohiuddin, et al., 1996           | Y                      | Y                  | Y                  | Y             | Y                       | Y                                | Y                                      | Y                        | Y                            | Y                                 | Y                                     | U                          | N  | N                            |
| Morise, et al., 2000              | Y                      | Y                  | Y                  | Y             | N                       | Y                                | Y                                      | Y                        | Y                            | Y                                 | Y                                     | Y                          | N  | N                            |
| Morise, et al., 1995-A            | U                      | Y                  | Y                  | N             | N                       | Y                                | Y                                      | Y                        | Y                            | U                                 | U                                     | U                          | N  | N                            |
| Morise, et al., 1995-B            | U                      | N                  | Y                  | Y             | Y                       | Y                                | Y                                      | N                        | Y                            | Y                                 | Y                                     | Y                          | N  | N                            |
| Morise, et al., 2002              | Y                      | Y                  | Y                  | Y             | Y                       | Y                                | Y                                      | Y                        | U                            | Y                                 | Y                                     | N                          | N  | N                            |
| Morise, et al., 2004 <sup>e</sup> | U                      | Y                  | Y                  | U             | U                       | Y                                | U                                      | U                        | U                            | U                                 | U                                     | U                          | Y  | U                            |
| Nascimento, et al., 2008          | Y                      | Y                  | Y                  | N             | N                       | Y                                | Y                                      | N                        | Y                            | Y                                 | Y                                     | U                          | N  | N                            |

| Study                       | Patient Representation | Selection Criteria | Reference Standard | Time Interval | Sample for Verification | Reference test uniformly applied | Reference test independently performed | Index Test Replicability | Reference Test Replicability | Index Test Blinded Interpretation | Reference Test Blinded Interpretation | Clinical Data Availability | Uninterpretable or Intermediate Test Reporting | Study Withdrawal Description |
|-----------------------------|------------------------|--------------------|--------------------|---------------|-------------------------|----------------------------------|--|--------------------------|------------------------------|-----------------------------------|---------------------------------------|----------------------------|--|------------------------------|
| Ozdemir, et al., 2002       | Y                      | Y                  | Y                  | U             | Y                       | Y                                | Y                                      | Y                        | Y                            | Y                                 | U                                     | U                          | N  | N                            |
| Pundziute, et al., 2008     | U                      | Y                  | Y                  | Y             | Y                       | Y                                | Y                                      | Y                        | U                            | Y                                 | Y                                     | U                          | Y  | Y                            |
| Raman, et al., 2008         | U                      | Y                  | Y                  | U             | N                       | U                                | U                                      | Y                        | U                            | Y                                 | U                                     | U                          | Y  | N                            |
| Richardson, et al., 1995    | N                      | Y                  | Y                  | Y             | Y                       | Y                                | Y                                      | Y                        | Y                            | U                                 | U                                     | U                          | N  | N                            |
| Robert, et al., 1991        | U                      | Y                  | Y                  | U             | Y                       | Y                                | U                                      | Y                        | U                            | U                                 | U                                     | U                          | N  | N                            |
| Roger, et al., 1997         | U                      | N                  | Y                  | N             | N                       | Y                                | Y                                      | Y                        | Y                            | Y                                 | Y                                     | U                          | N  | N                            |
| Rollan, et al., 2002        | U                      | Y                  | Y                  | U             | Y                       | Y                                | U                                      | U                        | U                            | U                                 | U                                     | U                          | N  | N                            |
| San Roman, et al., 1998     | U                      | Y                  | Y                  | Y             | Y                       | Y                                | Y                                      | Y                        | U                            | Y                                 | Y                                     | N                          | Y  | N                            |
| Sanfilippo, et al., 2005    | Y                      | Y                  | U                  | U             | U                       | N                                | N                                      | Y                        | N                            | Y                                 | U                                     | N                          | Y  | N                            |
| Santana-Boado, et al., 1998 | U                      | N                  | Y                  | Y             | Y                       | Y                                | Y                                      | Y                        | Y                            | Y                                 | Y                                     | Y                          | N  | N                            |
| Sawada, et al., 1989        | U                      | Y                  | Y                  | Y             | Y                       | Y                                | Y                                      | Y                        | U                            | Y                                 | Y                                     | U                          | Y  | N                            |
| Schupbach, et al., 2008     | U                      | N                  | Y                  | Y             | Y                       | Y                                | Y                                      | Y                        | U                            | Y                                 | Y                                     | U                          | N  | N                            |

| Study                           | Patient Representation | Selection Criteria | Reference Standard | Time Interval | Sample for Verification | Reference test uniformly applied | Reference test independently performed | Index Test Replicability | Reference Test Replicability | Index Test Blinded Interpretation | Reference Test Blinded Interpretation | Clinical Data Availability | Uninterpretable or Intermediate Test Reporting | Study Withdrawal Description |
|---------------------------------|------------------------|--------------------|--------------------|---------------|-------------------------|----------------------------------|--|--------------------------|------------------------------|-----------------------------------|---------------------------------------|----------------------------|--|------------------------------|
| Severi, et al., 1994            | Y                      | Y                  | Y                  | Y             | Y                       | Y                                | Y                                      | Y                        | Y                            | Y                                 | Y                                     | Y                          | Y  | N                            |
| Sharir, et al., 2000            | U                      | N                  | Y                  | U             | Y                       | Y                                | Y                                      | Y                        | U                            | U                                 | U                                     | U                          | N  | N                            |
| Shaw, et al., 2011 <sup>f</sup> | N                      | Y                  | U                  | Y             | N                       | N                                | N                                      | Y                        | Y                            | Y                                 | U                                     | U                          | U  | Y                            |
| Shi, et al., 2007               | U                      | N                  | Y                  | Y             | U                       | Y                                | U                                      | Y                        | N                            | U                                 | U                                     | U                          | U  | N                            |
| Shin, et al., 2003              | U                      | Y                  | Y                  | Y             | Y                       | Y                                | Y                                      | Y                        | U                            | Y                                 | U                                     | N                          | N  | N                            |
| Shivalkar, et al., 2007         | U                      | Y                  | Y                  | Y             | Y                       | Y                                | U                                      | Y                        | Y                            | U                                 | U                                     | U                          | Y  | U                            |
| Siegler, et al., 2011           | Y                      | Y                  | U                  | Y             | Y                       | Y                                | Y                                      | Y                        | Y                            | Y                                 | Y                                     | U                          | U  | N                            |
| Sinha, et al., 2008             | Y                      | Y                  | Y                  | Y             | Y                       | Y                                | Y                                      | N                        | N                            | U                                 | U                                     | U                          | N  | N                            |
| Sketch, et al., 1975            | U                      | N                  | Y                  | U             | Y                       | Y                                | U                                      | Y                        | U                            | U                                 | U                                     | U                          | U  | N                            |
| Slavich, et al., 1996           | U                      | Y                  | Y                  | Y             | Y                       | Y                                | U                                      | Y                        | U                            | Y                                 | U                                     | U                          | N  | N                            |
| Slomka, et al., 2007            | U                      | N                  | Y                  | Y             | Y                       | Y                                | Y                                      | Y                        | U                            | U                                 | U                                     | U                          | N  | N                            |
| Slomka, et al., 2005            | U                      | N                  | Y                  | Y             | Y                       | Y                                | Y                                      | Y                        | U                            | Y                                 | Y                                     | U                          | U  | N                            |

| Study                   | Patient Representation | Selection Criteria | Reference Standard | Time Interval | Sample for Verification | Reference test uniformly applied | Reference test independently performed | Index Test Replicability | Reference Test Replicability | Index Test Blinded Interpretation | Reference Test Blinded Interpretation | Clinical Data Availability | Uninterpretable or Intermediate Test Reporting | Study Withdrawal Description |
|-------------------------|------------------------|--------------------|--------------------|---------------|-------------------------|----------------------------------|--|--------------------------|------------------------------|-----------------------------------|---------------------------------------|----------------------------|--|------------------------------|
| Svart, et al., 2010     | U                      | N                  | Y                  | N             | Y                       | Y                                | Y                                      | Y                        | Y                            | U                                 | U                                     | U                          | N  | N                            |
| Taillefer, et al., 1997 | U                      | N                  | Y                  | U             | Y                       | Y                                | Y                                      | Y                        | Y                            | Y                                 | Y                                     | N                          | N  | N                            |
| Takeuchi, et al., 1996  | U                      | Y                  | Y                  | Y             | Y                       | Y                                | Y                                      | Y                        | U                            | Y                                 | Y                                     | U                          | Y  | Y                            |
| Travin, et al., 2000    | U                      | U                  | Y                  | U             | Y                       | Y                                | Y                                      | Y                        | U                            | Y                                 | Y                                     | U                          | Y  | N                            |
| Van Train, et al., 1994 | U                      | N                  | Y                  | U             | Y                       | Y                                | Y                                      | Y                        | Y                            | Y                                 | Y                                     | U                          | Y  | N                            |
| Vashist, et al., 2007   | U                      | Y                  | Y                  | Y             | Y                       | Y                                | U                                      | U                        | U                            | U                                 | U                                     | U                          | U  | N                            |
| Weiner, et al., 1985    | N                      | N                  | Y                  | Y             | Y                       | Y                                | Y                                      | Y                        | Y                            | Y                                 | Y                                     | U                          | Y  | N                            |
| Weiner, et al., 1979    | U                      | N                  | Y                  | Y             | Y                       | Y                                | Y                                      | Y                        | Y                            | U                                 | U                                     | U                          | N  | N                            |
| Weustink, et al., 2007  | N                      | Y                  | Y                  | U             | Y                       | Y                                | Y                                      | Y                        | U                            | Y                                 | Y                                     | U                          | Y  | N                            |
| Weustink, et al., 2009  | Y                      | Y                  | Y                  | U             | Y                       | Y                                | Y                                      | Y                        | Y                            | Y                                 | Y                                     | U                          | N  | N                            |
| Williams, et al., 1994  | Y                      | N                  | Y                  | U             | Y                       | Y                                | Y                                      | Y                        | Y                            | Y                                 | Y                                     | U                          | Y  | N                            |
| Wolak, et al., 2008     | U                      | Y                  | Y                  | Y             | Y                       | Y                                | U                                      | Y                        | U                            | Y                                 | U                                     | U                          | U  | N                            |



| Study                              | Patient Representation | Selection Criteria | Reference Standard | Time Interval | Sample for Verification | Reference test uniformly applied | Reference test independently performed | Index Test Replicability | Reference Test Replicability | Index Test Blinded Interpretation | Reference Test Blinded Interpretation | Clinical Data Availability | Uninterpretable or Intermediate Test Reporting | Study Withdrawal Description |
|------------------------------------|------------------------|--------------------|--------------------|---------------|-------------------------|----------------------------------|--|--------------------------|------------------------------|-----------------------------------|---------------------------------------|----------------------------|--|------------------------------|
| Wong, et al., 2001                 | U                      | N                  | Y                  | U             | N                       | N                                | Y                                      | N                        | N                            | U                                 | U                                     | U                          | N  | N                            |
| Yamauchi, et al., 1985             | U                      | Y                  | Y                  | U             | Y                       | Y                                | Y                                      | Y                        | U                            | U                                 | Y                                     | U                          | N  | N                            |
| Yeih, et al., 2007                 | U                      | N                  | Y                  | Y             | U                       | Y                                | Y                                      | Y                        | U                            | Y                                 | U                                     | U                          | N  | Y                            |
| <b>Totals</b>                      |                        |                    |                    |               |                         |                                  |  |                          |                              |                                   |                                       |                            |  |                              |
| <b>Y = Yes (low risk for bias)</b> | 36                     | 66                 | 100                | 65            | 84                      | 96                               | 81                                     | 91                       | 56                           | 71                                | 56                                    | 20                         | 34   | 22                           |
| <b>N = No (high risk for bias)</b> | 10                     | 35                 | 1                  | 7             | 15                      | 7                                | 4                                      | 7                        | 11                           | 3                                 | 3                                     | 12                         | 61   | 79                           |
| <b>U = Unclear</b>                 | 58                     | 3                  | 3                  | 32            | 5                       | 1                                | 19                                     | 6                        | 37                           | 30                                | 45                                    | 72                         | 9  | 3                            |

<sup>a</sup>Refer to Appendix B in this report for the 14 QUADAS questions. (For full details on QUADAS methodology, see: <http://www.biomedcentral.com/1471-2288/3/25>.)

<sup>b</sup>Related methods article: Dewey, et al., 2006 (refer to Appendix C for full citation).

<sup>c</sup>Related methods article: Miller, et al., 2009 (refer to Appendix C for full citation).

<sup>d</sup>Related methods articles: Meijboom, et al., 2007 and Meijboom, et al., 2008 (refer to Appendix C for full citations).

<sup>e</sup>Related methods article: Merz, et al., 1999 (refer to Appendix C for full citation).

<sup>f</sup>Related methods article: Mieres, et al., 2009 (refer to Appendix C for full citation).

Abbreviations: N = No; U = Unclear, Y = Yes

## **Appendix E. List of Excluded Studies**

All studies listed below were reviewed in their full-text version and excluded. Following each reference, in italics, is the reason for exclusion. Reasons for exclusion signify only the usefulness of the articles for this study and are not intended as criticisms of the articles.

Ababneh AA, Sciacca RR, Kim B,, et al. Normal limits for left ventricular ejection fraction and volumes estimated with gated myocardial perfusion imaging in patients with normal exercise test results: influence of tracer, gender, and acquisition camera. *J Nucl Cardiol* 2000;7(6):661-8. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; No outcomes of interest.*

Abaci A, Oguzhan A, Topsakal R,, et al. Intracoronary electrocardiogram and angina pectoris during percutaneous coronary interventions as an assessment of myocardial viability: comparison with low-dose dobutamine echocardiography. *Catheter Cardiovasc Interv* 2003;60(4):469-76. *Full-text exclusion reason(s): No data for NITs of interest (ECG, ECHO, SPECT, PET, CMR, CTA).*

Abbara S, Chow BJ, Pena AJ,, et al. Assessment of left ventricular function with 16- and 64-slice multi-detector computed tomography. *Eur J Radiol* 2008;67(3):481-6. *Full-text exclusion reason(s): Data for women not reported as a subgroup; No outcomes of interest.*

Abbott BG, Abdel-Aziz I, Nagula S,, et al. Selective use of single-photon emission computed tomography myocardial perfusion imaging in a chest pain center. *Am J Cardiol* 2001;87(12):1351-5. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Abbott BG, Afshar M, Berger AK,, et al. Prognostic significance of ischemic electrocardiographic changes during adenosine infusion in patients with normal myocardial perfusion imaging. *J Nucl Cardiol* 2003;10(1):9-16. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Abdelmoneim SS, Bernier M, Dhoble A,, et al. Assessment of myocardial perfusion during adenosine stress using real time three-dimensional and two-dimensional myocardial contrast echocardiography: comparison with single-photon emission computed tomography. *Echocardiography* 2010;27(4):421-9. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Abdelmoneim SS, Bernier M, Dhoble A,, et al. Diagnostic accuracy of contrast echocardiography during adenosine stress for detection of abnormal myocardial perfusion: a prospective comparison with technetium-99 m sestamibi single-photon emission computed tomography. *Heart Vessels* 2010;25(2):121-30. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Abdelmoneim SS, Bernier M, Scott CG,, et al. Safety of contrast agent use during stress echocardiography: a 4-year experience from a single-center cohort study of 26,774 patients. *JACC Cardiovasc Imaging* 2009;2(9):1048-56. *Full-text exclusion reason(s): No data for NITs of interest (ECG, ECHO, SPECT, PET, CMR, CTA).*

Abdelmoneim SS, Dhoble A, Bernier M,, et al. Quantitative myocardial contrast echocardiography during pharmacological stress for diagnosis of coronary artery disease: a systematic review and meta-analysis of diagnostic accuracy studies. *Eur J Echocardiogr* 2009;10(7):813-25. *Full-text exclusion reason(s): Not a clinical study report.*

Abdelmoneim SS, Dhoble A, Bernier M,, et al. Absolute myocardial blood flow determination using real-time myocardial contrast echocardiography during adenosine stress: comparison with single-photon emission computed tomography. *Heart* 2009;95(20):1662-8. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Abdullah SM, Khera A, Das SR,, et al. Relation of coronary atherosclerosis determined by electron beam computed tomography and plasma levels of n-terminal pro-brain natriuretic peptide in a multiethnic population-based sample (the Dallas Heart Study). *Am J Cardiol* 2005;96(9):1284-9. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup; No outcomes of interest.*

Abidov A, Bax JJ, Hayes SW,, et al. Transient ischemic dilation ratio of the left ventricle is a significant predictor of future cardiac events in patients with otherwise normal myocardial perfusion SPECT. *J Am Coll Cardiol* 2003;42(10):1818-25. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Abidov A, Gallagher MJ, Chinnaiyan KM,, et al. Clinical effectiveness of coronary computed tomographic angiography in the triage of patients to cardiac catheterization and revascularization after inconclusive stress testing: results of a 2-year prospective trial. *J Nucl Cardiol* 2009;16(5):701-13. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Abitbol E, Monin JL, Garot J,, et al. Relationship between the ischemic threshold at the onset of wall-motion abnormality on semisupine exercise echocardiography and the extent of coronary artery disease. *J Am Soc Echocardiogr* 2004;17(2):121-5. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Aboul-Enein FA, Hayes SW, Matsumoto N,, et al. Rest perfusion defects in patients with no history of myocardial infarction predict the presence of a critical coronary artery stenosis. *J Nucl Cardiol* 2003;10(6):656-62. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Abraham A, Kass M, Ruddy TD,, et al. Right and left ventricular uptake with Rb-82 PET myocardial perfusion imaging: markers of left main or 3 vessel disease. *J Nucl Cardiol* 2010;17(1):52-60. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Acampa W, Petretta M, Florimonte L,, et al. Prognostic value of exercise cardiac tomography performed late after percutaneous coronary intervention in symptomatic and symptom-free patients. *Am J Cardiol* 2003;91(3):259-63. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Acarturk E, Bozkurt A, Cayli M,, et al. Mitral annular calcification and aortic valve calcification may help in predicting significant coronary artery disease. *Angiology* 2003;54(5):561-7. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Achenbach S, Giesler T, Ropers D,, et al. Detection of coronary artery stenoses by contrast-enhanced, retrospectively electrocardiographically-gated, multislice spiral computed tomography. *Circulation* 2001;103(21):2535-8. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Achenbach S, Goroll T, Seltmann M,, et al. Detection of coronary artery stenoses by low-dose, prospectively ECG-triggered, high-pitch spiral coronary CT angiography. *JACC Cardiovasc Imaging* 2011;4(4):328-37. *Full-text exclusion reason(s): Data for women not reported as a subgroup; No outcomes of interest.*

Achenbach S, Ropers D, Pohle FK,, et al. Detection of coronary artery stenoses using multi-detector CT with 16 x 0.75 collimation and 375 ms rotation. *Eur Heart J* 2005;26(19):1978-86. *Full-text exclusion reason(s): Data for women not reported as a subgroup; No outcomes of interest.*

Achenbach S, Ropers U, Kuettner A,, et al. Randomized comparison of 64-slice single- and dual-source computed tomography coronary angiography for the detection of coronary artery disease. *JACC Cardiovasc Imaging* 2008;1(2):177-86. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Adams GL, Trimble MA, Brosnan RB,, et al. Evaluation of combined cardiac positron emission tomography and coronary computed tomography angiography for the detection of coronary artery disease. *Nucl Med Commun* 2008;29(7):593-8. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Adamu U, Knollmann D, Almutairi B,, et al. Stress/rest myocardial perfusion scintigraphy in patients without significant coronary artery disease. *J Nucl Cardiol* 2010;17(1):38-44. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Adler Y, Vaturi M, Herz I,, et al. Nonobstructive aortic valve calcification: a window to significant coronary artery disease. *Atherosclerosis* 2002;161(1):193-7. *Full-text exclusion reason(s): No data for NITs of interest (ECG, ECHO, SPECT, PET, CMR, CTA).*

Aepfelbacher FC, Johnson RB, Schwartz JG,, et al. Validation of a model of left ventricular segmentation for interpretation of SPET myocardial perfusion images. *Eur J Nucl Med* 2001;28(11):1624-9. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Aessopos A, Tsironi M, Vassiliadis I, et al. Exercise-induced myocardial perfusion abnormalities in sickle beta-thalassemia: Tc-99m tetrofosmin gated SPECT imaging study. *Am J Med* 2001;111(5):355-60. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; No outcomes of interest.*

Afonso L, Mahajan N. Single-photon emission computed tomography myocardial perfusion imaging in the diagnosis of left main disease. *Clin Cardiol* 2009;32(12):E11-5. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD.*

Agarwal PP, Patel S, Corbett J, et al. Left ventricular functional analysis with 16- and 64-row multidetector computed tomography: comparison with gated single-photon emission computed tomography. *J Comput Assist Tomogr* 2009;33(1):8-14. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup; No outcomes of interest.*

Aggeli C, Christoforatu E, Giannopoulos G, et al. The diagnostic value of adenosine stress-contrast echocardiography for diagnosis of coronary artery disease in hypertensive patients: comparison to Tl-201 single-photon emission computed tomography. *Am J Hypertens* 2007;20(5):533-8. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup.*

Aggeli C, Giannopoulos G, Misovoulos P, et al. Real-time three-dimensional dobutamine stress echocardiography for coronary artery disease diagnosis: validation with coronary angiography. *Heart* 2007;93(6):672-5. *Full-text exclusion reason(s): Data for women not reported as a subgroup; No outcomes of interest.*

Ahlberg AW, Baghdasarian SB, Athar H, et al. Symptom-limited exercise combined with dipyridamole stress: prognostic value in assessment of known or suspected coronary artery disease by use of gated SPECT imaging. *J Nucl Cardiol* 2008;15(1):42-56. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; No outcomes of interest.*

Ahmad M, Xie T, McCulloch M, et al. Real-time three-dimensional dobutamine stress echocardiography in assessment stress echocardiography in assessment of ischemia: comparison with two-dimensional dobutamine stress echocardiography. *J Am Coll Cardiol* 2001;37(5):1303-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Ahmadi N, Nabavi V, Hajsadeghi F, et al. Impaired aortic distensibility measured by computed tomography is associated with the severity of coronary artery disease. *Int J Cardiovasc Imaging* 2010. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Study does not compare one NIT to another or to diagnostic cardiac catheterization; No outcomes of interest.*

Ahmadi N, Nabavi V, Hajsadeghi F, et al. Mortality incidence of patients with non-obstructive coronary artery disease diagnosed by computed tomography angiography. *Am J Cardiol* 2011;107(1):10-6. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Ahmadi N, Shavelle D, Nabavi V, et al. Coronary distensibility index measured by computed tomography is associated with the severity of coronary artery disease. *J Cardiovasc Comput Tomogr* 2010;4(2):119-26. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup; No outcomes of interest.*

Ahmed B, Merz CNB, Sopko G. Are we 'WISE'r? Findings from the NHLBI-sponsored Women's Ischemia Syndrome Evaluation study. *Women's Health* 2006;2(1):57-64. *Full-text exclusion reason(s): Not a clinical study report.*

Ahn JM, Kang SJ, Mintz GS, et al. Validation of minimal luminal area measured by intravascular ultrasound for assessment of functionally significant coronary stenosis: Comparison with myocardial perfusion imaging. *JACC: Cardiovascular Interventions* 2011;4(6):665-671. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Akinboboye OO, Idris O, Onwuanyi A, et al. Incidence of major cardiovascular events in black patients with normal myocardial stress perfusion study results. *J Nucl Cardiol* 2001;8(5):541-7. *Full-text exclusion reason(s): No*

women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup; No outcomes of interest.

Akram K, Voros S. Absolute coronary artery calcium scores are superior to MESA percentile rank in predicting obstructive coronary artery disease. *Int J Cardiovasc Imaging* 2008;24(7):743-9. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Study does not compare one NIT to another or to diagnostic cardiac catheterization.*

Al Moudi M, Sun Z, Lenzo N. Diagnostic value of SPECT, PET and PET/CT in the diagnosis of coronary artery disease: A systematic review. *Biomedical Imaging and Intervention Journal* 2011;7(2). *Full-text exclusion reason(s): Conference abstract or trial registry posting.*

Al-Attar AT, Mahussain SA, Sadanandan S. Cardiac tests in asymptomatic type 2 diabetics. *Med Princ Pract* 2002;11(4):171-5. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Albro PC, Gould KL, Westcott RJ, et al. Noninvasive assessment of coronary stenoses by myocardial imaging during pharmacologic coronary vasodilatation. III. Clinical trial. *Am J Cardiol* 1978;42(5):751-60. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Aldrovandi A, Cademartiri F, Menozzi A, et al. Evaluation of coronary atherosclerosis by multislice computed tomography in patients with acute myocardial infarction and without significant coronary artery stenosis: a comparative study with quantitative coronary angiography. *Circ Cardiovasc Imaging* 2008;1(3):205-11. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Aldrovandi A, Maffei E, Palumbo A, et al. Prognostic value of computed tomography coronary angiography in patients with suspected coronary artery disease: a 24-month follow-up study. *Eur Radiol* 2009;19(7):1653-60. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Alessandri N, Di Matteo A, Rondoni G, et al. Heart imaging: the accuracy of the 64-MSCT in the detection of coronary artery disease. *Eur Rev Med Pharmacol Sci* 2009;13(3):163-71. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Alexander W, Mehta SR, Giugliano RP. American Heart Association 2008 Scientific Sessions: The TMACS trial. *P and T* 2009;34(1):30. *Full-text exclusion reason(s): Conference abstract or trial registry posting.*

Alexanderson E, Mannting F, Gomez-Martin D, et al. Technetium-99m-Sestamibi SPECT myocardial perfusion imaging in patients with complete left bundle branch block. *Arch Med Res* 2004;35(2):150-6. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Alexopoulos D, Stathopoulos C, Kotrsaridis A, et al. Coronary artery calcium by digital cinefluoroscopy in patients with pain suggestive of an acute coronary syndrome. *Clin Cardiol* 2005;28(2):81-4. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Ali M, Mallick NH, Abid AR, et al. Significance of perfusion defects on dipyridamole thallium cardiac SPECT in patients with left bundle branch block. *J Ayub Med Coll Abbottabad* 2007;19(4):21-5. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Alkadhi H, Scheffel H, Desbiolles L, et al. Dual-source computed tomography coronary angiography: influence of obesity, calcium load, and heart rate on diagnostic accuracy. *Eur Heart J* 2008;29(6):766-76. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Alkadhi H, Stolzmann P, Desbiolles L, et al. Low-dose, 128-slice, dual-source CT coronary angiography: accuracy and radiation dose of the high-pitch and the step-and-shoot mode. *Heart* 2010;96(12):933-8. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Alkadhi H, Stolzmann P, Desbiolles L, et al. Low-dose, 128-slice, dual-source CT coronary angiography: accuracy and radiation dose of the high-pitch and the step-and-shoot mode. *Heart*; 2010:933-8.

Al-Khalili F, Janszky I, Andersson A., et al. Physical activity and exercise performance predict long-term prognosis in middle-aged women surviving acute coronary syndrome. *J Intern Med* 2007;261(2):178-87. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; No outcomes of interest.*

Allison JS, Heo J, Iskandrian AE. Artificial neural network modeling of stress single-photon emission computed tomographic imaging for detecting extensive coronary artery disease. *Am J Cardiol* 2005;95(2):178-81. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Allman KC, Berry J, Sucharski LA., et al. Determination of extent and location of coronary artery disease in patients without prior myocardial infarction by thallium-201 tomography with pharmacologic stress. *J Nucl Med* 1992;33(12):2067-73. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Al-Mallah M, Alqaisi F, Arafeh A., et al. Long term favorable prognostic value of negative treadmill echocardiogram in the setting of abnormal treadmill electrocardiogram: a 95 month median duration follow-up study. *J Am Soc Echocardiogr* 2008;21(9):1018-22. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Al-Mallah MH, Hachamovitch R, Dorbala S., et al. Incremental prognostic value of myocardial perfusion imaging in patients referred to stress single-photon emission computed tomography with renal dysfunction. *Circ Cardiovasc Imaging* 2009;2(6):429-36. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Al-Mallah MH, Sitek A, Moore SC., et al. Assessment of myocardial perfusion and function with PET and PET/CT. *J Nucl Cardiol* 2010;17(3):498-513. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Almeda FQ, Shah R, Senter S., et al. Clinical and angiographic profile of patients with markedly elevated coronary calcium scores ( $\geq 1000$ ) detected by electron beam computed tomography. *Cardiovasc Radiat Med* 2004;5(3):109-12. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Almeida AG, Gabriel HM, Coutinho CA., et al. Myocardial perfusion and angioplasty. Comparison of myocardial contrast echocardiography and scintigraphy. *Rev Port Cardiol* 2002;21(7-8):859-68. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Almeida MC, Markman Filho B. Prognostic value of dipyridamole stress echocardiography in women. *Arq Bras Cardiol* 2011;96(1):31-7. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization.*

Alqaisi F, Albadarin F, Jaffery Z., et al. Prognostic predictors and outcomes in patients with abnormal myocardial perfusion imaging and angiographically insignificant coronary artery disease. *J Nucl Cardiol* 2008;15(6):754-61. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Al-Saadi N, Nagel E, Gross M., et al. Noninvasive detection of myocardial ischemia from perfusion reserve based on cardiovascular magnetic resonance. *Circulation* 2000;101(12):1379-83. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Amanullah AM. Diagnostic and prognostic value of myocardial perfusion imaging in patients with known or suspected stable coronary artery disease. *Echocardiography* 2000;17(6 Pt 1):587-95. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Amanullah AM, Berman DS, Kiat H., et al. Usefulness of hemodynamic changes during adenosine infusion in predicting the diagnostic accuracy of adenosine technetium-99m sestamibi single-photon emission computed tomography (SPECT). *Am J Cardiol* 1997;79(10):1319-22. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Amanullah AM, Bevegard S, Lindvall K,, et al. Assessment of left ventricular wall motion in angina pectoris by two-dimensional echocardiography and myocardial perfusion by technetium-99m sestamibi tomography during adenosine-induced coronary vasodilation and comparison with coronary angiography. *Am J Cardiol* 1993;72(14):983-9. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Ambrosi P, Bruneau P, Faugere G,, et al. [Prognostic value of thallium-201 myocardial scintigraphy in patients with hypertension, suspected of coronary disease]. *Ann Cardiol Angeiol (Paris)* 1993;42(9):479-83. *Full-text exclusion reason(s): Non-English.*

Amemiya S, Takao H. Computed tomographic coronary angiography for diagnosing stable coronary artery disease: a cost-utility and cost-effectiveness analysis. *Circ J* 2009;73(7):1263-70. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Amici E, Cortigiani L, Coletta C,, et al. Usefulness of pharmacologic stress echocardiography for the long-term prognostic assessment of patients with typical versus atypical chest pain. *Am J Cardiol* 2003;91(4):440-2. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Ammann P, Naegeli B, Rickli H,, et al. Characteristics of patients with abnormal stress technetium Tc 99m sestamibi SPECT studies without significant coronary artery diameter stenoses. *Clin Cardiol* 2003;26(11):521-4. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup; No outcomes of interest.*

Anagnostopoulos C, Almonacid A, El Fakhri G,, et al. Quantitative relationship between coronary vasodilator reserve assessed by 82Rb PET imaging and coronary artery stenosis severity. *Eur J Nucl Med Mol Imaging* 2008;35(9):1593-601. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup; No outcomes of interest.*

Anand DV, Lim E, Raval U,, et al. Prevalence of silent myocardial ischemia in asymptomatic individuals with subclinical atherosclerosis detected by electron beam tomography. *J Nucl Cardiol* 2004;11(4):450-7. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Andersen K, Hennersdorf M, Cohnen M,, et al. Myocardial delayed contrast enhancement in patients with arterial hypertension: initial results of cardiac MRI. *Eur J Radiol* 2009;71(1):75-81. *Full-text exclusion reason(s): Data for women not reported as a subgroup; No outcomes of interest.*

Andrade JM, Gowdak LH, Giorgi MC,, et al. Cardiac MRI for detection of unrecognized myocardial infarction in patients with end-stage renal disease: comparison with ECG and scintigraphy. *AJR Am J Roentgenol* 2009;193(1):W25-32. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup; No outcomes of interest.*

Andrade MJ, Picano E, Pingitore A,, et al. Dipyridamole stress echocardiography in patients with severe left main coronary artery narrowing. Echo Persantine International Cooperative (EPIC) Study Group—Subproject "Left Main Detection". *Am J Cardiol* 1994;73(7):450-5. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Andreini D, Pontone G, Ballerini G,, et al. Feasibility and diagnostic accuracy of 16-slice multidetector computed tomography coronary angiography in 500 consecutive patients: critical role of heart rate. *Int J Cardiovasc Imaging* 2007;23(6):789-801. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*



Andreini D, Pontone G, Bartorelli AL, et al. Comparison of the diagnostic performance of 64-slice computed tomography coronary angiography in diabetic and non-diabetic patients with suspected coronary artery disease. *Cardiovasc Diabetol* 2010;9:80. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Anjaneyulu A, Raghu K, Chandramukhi S, et al. Evaluation of left main coronary artery stenosis by transthoracic echocardiography. *J Am Soc Echocardiogr* 2008;21(7):855-60. *Full-text exclusion reason(s): No data for NITs of interest (ECG, ECHO, SPECT, PET, CMR, CTA).*

Anthopoulos LP, Bonou MS, Kardaras FG, et al. Stress echocardiography in elderly patients with coronary artery disease: applicability, safety and prognostic value of dobutamine and adenosine echocardiography in elderly patients. *J Am Coll Cardiol* 1996;28(1):52-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Anthopoulos LP, Bonou MS, Sioras EP, et al. Echocardiographic detection of the extent of coronary artery disease in the elderly using dobutamine and adenosine infusion. *Coron Artery Dis* 1997;8(10):633-43. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Arantes RL, Gowdak LH, Paula FJ, et al. Myocardial scintigraphy and clinical stratification as predictors of events in renal transplant candidates. *J Nephrol* 2010;23(3):314-20. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Ariff B, Thom SA, Foale RA, et al. Stress echocardiography for the diagnosis of ischaemia in hypertensives. *Journal of human hypertension* 2000(6):399-401. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Ariyaratajah V, Kranis M, Apiyasawat S, et al. Association of myocardial ischemia and coronary angiographic lesions with increased left atrial dimension during exercise tolerance tests among patients without known coronary heart disease. *Am J Cardiol* 2007;99(9):1187-92. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Arnold J, Selvanayagam J. Non-invasive Detection of Coronary Artery Disease Using Blood Oxygen Level-dependent Magnetic Resonance Imaging at 3 Tesla. 2009 American Heart Association Scientific Sessions, Orange County Convention Center, Orlando, Florida, 14-18 Nov 2009 2009. *Full-text exclusion reason(s): Conference abstract or trial registry posting.*

Arnold JR, Karamitsos TD, Bhamra-Ariza P, et al. Blood oxygen level-dependent magnetic resonance imaging at 3 Tesla in coronary artery disease: Validation using quantitative coronary angiography and cardiovascular magnetic resonance perfusion imaging. *Journal of Cardiovascular Magnetic Resonance* 2010;12:35. *Full-text exclusion reason(s): Conference abstract or trial registry posting*

Arnoldi E, Gebregziabher M, Schoepf UJ, et al. Automated computer-aided stenosis detection at coronary CT angiography: initial experience. *Eur Radiol* 2010;20(5):1160-7. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Arruda AL, Barretto RB, Shub C, et al. Prognostic significance of ST-segment elevation during dobutamine stress echocardiography. *Am Heart J* 2006;151(3):744 e1-744 e6. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Arruda AM, Das MK, Roger VL, et al. Prognostic value of exercise echocardiography in 2,632 patients > or = 65 years of age. *J Am Coll Cardiol* 2001;37(4):1036-41. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Arruda-Olson AM, Juracan EM, Mahoney DW, et al. Prognostic value of exercise echocardiography in 5,798 patients: is there a gender difference? *J Am Coll Cardiol* 2002;39(4):625-31. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization.*

Ashley EA, Myers J, Froelicher V. Exercise testing in clinical medicine. *Lancet* 2000;356(9241):1592-1597. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Askew JW, Miller TD, Araoz PA, et al. Abnormal electron beam computed tomography results: the value of repeating myocardial perfusion single-photon emission computed tomography in the ongoing assessment of coronary artery disease. *Mayo Clin Proc* 2008;83(1):17-22. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Aslam M, Zia ul S, Anwar MA, et al. Accuracy of thallium scintigraphy versus coronary angiography for coronary artery disease in diabetics. *J Coll Physicians Surg Pak* 2010;20(1):6-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Assomull RG, Lyne JC, Keenan N, et al. The role of cardiovascular magnetic resonance in patients presenting with chest pain, raised troponin, and unobstructed coronary arteries. *Eur Heart J* 2007;28(10):1242-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup; No outcomes of interest.*

Astarita C, Palinkas A, Nicolai E, et al. Dipyridamole-atropine stress echocardiography versus exercise SPECT scintigraphy for detection of coronary artery disease in hypertensives with positive exercise test. *J Hypertens* 2001;19(3):495-502. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Astorri E, Pattoneri P, Calbiani B, et al. Thallium-201 myocardial perfusion imaging in patients with systemic lupus erythematosus. *Minerva Cardioangiol* 2004;52(1):49-54. *Full-text exclusion reason(s): Not a clinical study report.*

Atak R, Turhan H, Sezgin AT, et al. Effects of slow coronary artery flow on QT interval duration and dispersion. *Ann Noninvasive Electrocardiol* 2003;8(2):107-11. *Full-text exclusion reason(s): No data for NITs of interest (ECG, ECHO, SPECT, PET, CMR, CTA).*

Atar S, Cercek B, Nagai T, et al. Transthoracic stress echocardiography with transesophageal atrial pacing for bedside evaluation of inducible myocardial ischemia in patients with new-onset chest pain. *Am J Cardiol* 2000;86(1):12-6. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Atar S, Feldman A, Darawshe A, et al. Utility and diagnostic accuracy of hand-carried ultrasound for emergency room evaluation of chest pain. *Am J Cardiol* 2004;94(3):408-9. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup; No outcomes of interest.*

Atar S, Nagai T, Cercek B, et al. Pacing stress echocardiography: an alternative to pharmacologic stress testing. *J Am Coll Cardiol* 2000;36(6):1935-41. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Atchley AE, Iskandrian AE, Bensimhon D, et al. Relationship of technetium-99m tetrofosmin-gated rest single-photon emission computed tomography myocardial perfusion imaging to death and hospitalization in heart failure patients: Results from the nuclear ancillary study of the HF-ACTION trial. *Am Heart J* 2011;161(6):1038-1045. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Atsma DE, Bavelaar-Croon CD, Germano G, et al. Good correlation between gated single photon emission computed myocardial tomography and contrast ventriculography in the assessment of global and regional left ventricular function. *Int J Card Imaging* 2000;16(6):447-53. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup; No outcomes of interest.*

Attar MN, Wong K, Groves DG, et al. Clinical implications of QRS duration and QT peak prolongation in patients with suspected coronary disease referred for elective cardiac catheterization. *Ann Noninvasive Electrocardiol* 2008;13(2):106-12. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Auseon AJ, Advani SS, Bush CA,, et al. Impact of 64-slice multidetector computed tomography on other diagnostic studies for coronary artery disease. *Am J Med* 2009;122(4):387-91. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Auseon AJ, Tran T, Garcia AM,, et al. Aortic pathophysiology by cardiovascular magnetic resonance in patients with clinical suspicion of coronary artery disease. *J Cardiovasc Magn Reson* 2007;9(1):43-8. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup; No outcomes of interest.*

Aviram G, Finkelstein A, Herz I,, et al. Clinical value of 16-slice multi-detector CT compared to invasive coronary angiography. *Int J Cardiovasc Intervent* 2005;7(1):21-8. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Aydinalp A, Bal U, Atar I,, et al. Value of stress myocardial perfusion scanning in diagnosis of severe coronary artery disease in liver transplantation candidates. *Transplant Proc* 2009;41(9):3757-60. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Bacci S, Villella M, Villella A,, et al. Screening for silent myocardial ischaemia in type 2 diabetic patients with additional atherogenic risk factors: applicability and accuracy of the exercise stress test. *Eur J Endocrinol* 2002;147(5):649-54. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup.*

Bachar GN, Atar E, Fuchs S,, et al. Prevalence and clinical predictors of atherosclerotic coronary artery disease in asymptomatic patients undergoing coronary multidetector computed tomography. *Coron Artery Dis* 2007;18(5):353-60. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Badano LP, Muraru D, Rigo F,, et al. High volume-rate three-dimensional stress echocardiography to assess inducible myocardial ischemia: a feasibility study. *J Am Soc Echocardiogr* 2010;23(6):628-35. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Badruddin SM, Ahmad A, Mickelson J,, et al. Supine bicycle versus post-treadmill exercise echocardiography in the detection of myocardial ischemia: a randomized single-blind crossover trial. *J Am Coll Cardiol* 1999;33(6):1485-90. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Baghdasarian SB, Heller GV. The role of myocardial perfusion imaging in the diagnosis of patients with coronary artery disease: developments over the past year. *Curr Opin Cardiol* 2005;20(5):369-74. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Baik HK, Budoff MJ, Lane KL,, et al. Accurate measures of left ventricular ejection fraction using electron beam tomography: a comparison with radionuclide angiography, and cine angiography. *Int J Card Imaging* 2000;16(5):391-8. *Full-text exclusion reason(s): Data for women not reported as a subgroup; No outcomes of interest.*

Bailon R, Mateo J, Olmos S,, et al. Coronary artery disease diagnosis based on exercise electrocardiogram indexes from repolarisation, depolarisation and heart rate variability. *Med Biol Eng Comput* 2003;41(5):561-71. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Balaravi B, Miller TD, Hodge DO,, et al. The value of stress single photon emission computed tomography in patients without known coronary artery disease presenting with dyspnea. *Am Heart J* 2006;152(3):551-7. *Full-text exclusion reason(s): Data for women not reported as a subgroup; No outcomes of interest.*

Balnave K, Scott ME, Morton P,, et al. Reliable prediction of coronary disease using treadmill exercise testing. *Br Med J* 1978;1(6118):958-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

- Bamberg F, Becker A, Schwarz F, et al. Detection of hemodynamically significant coronary artery stenosis: incremental diagnostic value of dynamic CT-based myocardial perfusion imaging. *Radiology* 2011;260(3):689-98. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*
- Bamberg F, Sommer WH, Hoffmann V, et al. Meta-analysis and systematic review of the long-term predictive value of assessment of coronary atherosclerosis by contrast-enhanced coronary computed tomography angiography. *J Am Coll Cardiol* 2011;57(24):2426-36. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*
- Bamberg F, Truong QA, Blankstein R, et al. Usefulness of age and gender in the early triage of patients with acute chest pain having cardiac computed tomographic angiography. *Am J Cardiol* 2009;104(9):1165-70. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization.*
- Bampi AB, Rochitte CE, Favarato D, et al. Comparison of non-invasive methods for the detection of coronary atherosclerosis. *Clinics (Sao Paulo)* 2009;64(7):675-82. *Full-text exclusion reason(s): No data for NITs of interest (ECG, ECHO, SPECT, PET, CMR, CTA).*
- Banerjee SK, Haque KM, Sharma AK, et al. Role of exercise tolerance test (ETT) and gated single photon emission computed tomography-myocardial perfusion imaging (SPECT-MPI) in predicting severity of ischemia in patients with chest pain. *Bangladesh Med Res Counc Bull* 2005;31(1):27-35. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*
- Bangalore S, Gopinath D, Yao SS, et al. Risk stratification using stress echocardiography: incremental prognostic value over historic, clinical, and stress electrocardiographic variables across a wide spectrum of bayesian pretest probabilities for coronary artery disease. *J Am Soc Echocardiogr* 2007;20(3):244-52. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization.*
- Banzo I, Pena FJ, Allende RH, et al. Prospective clinical comparison of non-corrected and attenuation- and scatter-corrected myocardial perfusion SPECT in patients with suspicion of coronary artery disease. *Nucl Med Commun* 2003;24(9):995-1002. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*
- Barbirato GB, Azevedo JC, Felix RC, et al. Use of resting myocardial scintigraphy during chest pain to exclude diagnosis of acute myocardial infarction. *Arq Bras Cardiol* 2009;92(4):269-74. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*
- Barmeyer AA, Stork A, Muellerleile K, et al. Contrast-enhanced cardiac MR imaging in the detection of reduced coronary flow velocity reserve. *Radiology* 2007;243(2):377-85. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*
- Bart BA, Cen YY, Hendel RC, et al. Comparison of dobutamine stress echocardiography, dobutamine SPECT, and adenosine SPECT myocardial perfusion imaging in patients with end-stage renal disease. *J Nucl Cardiol* 2009;16(4):507-15. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*
- Bart BA, Erlien DA, Herzog CA, et al. Marked differences between patients referred for stress echocardiography and myocardial perfusion imaging studies. *Am Heart J* 2005;149(5):888-93. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup; No outcomes of interest.*
- Bartel T, Yang Y, Muller S, et al. Noninvasive assessment of microvascular function in arterial hypertension by transthoracic Doppler harmonic echocardiography. *J Am Coll Cardiol* 2002;39(12):2012-8. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*
- Basic D, Siu SC, Skyba DM, et al. Prognostic value of myocardial perfusion contrast echocardiography in patients with suggested or known ischemic heart disease. *J Am Soc Echocardiogr* 2006;19(10):1203-10. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*
- Baskot B, Obradovic S, Rafajlovski S, et al. Adenosine stress protocols for nuclear cardiology imaging. *Prilozi* 2008;29(1):281-9. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Study does not compare one*

*NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup; No outcomes of interest.*

Bastarrika G, Ramos-Duran L, Rosenblum MA, et al. Adenosine-stress dynamic myocardial CT perfusion imaging: initial clinical experience. *Invest Radiol* 2010;45(6):306-13. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Bateman TM, Heller GV, McGhie AI, et al. Multicenter investigation comparing a highly efficient half-time stress-only attenuation correction approach against standard rest-stress Tc-99m SPECT imaging. *J Nucl Cardiol* 2009;16(5):726-735. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Bateman TM, Heller GV, McGhie AI, et al. Diagnostic accuracy of rest/stress ECG-gated Rb-82 myocardial perfusion PET: comparison with ECG-gated Tc-99m sestamibi SPECT. *J Nucl Cardiol* 2006;13(1):24-33. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Battle E, Vilacosta I, San Roman JA, et al. [Elective noninvasive test in the diagnosis of coronary disease in the aged]. *Rev Esp Cardiol* 1998;51(1):35-42. *Full-text exclusion reason(s): Non-English.*

Bauer RW, Thilo C, Chiaramida SA, et al. Noncalcified atherosclerotic plaque burden at coronary CT angiography: a better predictor of ischemia at stress myocardial perfusion imaging than calcium score and stenosis severity. *AJR Am J Roentgenol* 2009;193(2):410-8. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Bauernfeind T, Preda I, Szakolczai K, et al. Diagnostic value of the left atrial electrical potentials detected by body surface potential mapping in the prediction of coronary artery disease. *Int J Cardiol* 2011;150(3):315-318. *Full-text exclusion reason(s): No data for NITs of interest (ECG, ECHO, SPECT, PET, CMR, CTA).*

Baumgart D, Schmermund A, Goerge G, et al. Comparison of electron beam computed tomography with intracoronary ultrasound and coronary angiography for detection of coronary atherosclerosis. *J Am Coll Cardiol* 1997;30(1):57-64. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Baumuller S, Leschka S, Desbiolles L, et al. Dual-source versus 64-section CT coronary angiography at lower heart rates: comparison of accuracy and radiation dose. *Radiology* 2009;253(1):56-64. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Bayrak F, Guneyusu T, Gemici G, et al. Diagnostic performance of 64-slice computed tomography coronary angiography to detect significant coronary artery stenosis. *Acta Cardiol* 2008;63(1):11-7. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Beanlands RS, Nichol G, Huszti E, et al. F-18-fluorodeoxyglucose positron emission tomography imaging-assisted management of patients with severe left ventricular dysfunction and suspected coronary disease: a randomized, controlled trial (PARR-2). *J Am Coll Cardiol* 2007;50(20):2002-12. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Beck T, Burgstahler C, Kuettner A, et al. Clinical use of multislice spiral computed tomography in 210 highly preselected patients: experience with 4 and 16 slice technology. *Heart* 2005;91(11):1423-7. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Beck T, Kuettner A, Burgstahler C, et al. [Noninvasive detection of coronary stenosis using 16-slice detector computed tomography in carefully selected patients]. *Med Klin (Munich)* 2004;99(11):645-50. *Full-text exclusion reason(s): Non-English.*

Becker A, Leber A, Becker C, et al. Predictive value of coronary calcifications for future cardiac events in asymptomatic individuals. *Am Heart J* 2008;155(1):154-60. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Study does not compare one NIT to another or to diagnostic cardiac catheterization.*

Becker A, Leber A, White CW, et al. Multislice computed tomography for determination of coronary artery disease in a symptomatic patient population. *Int J Cardiovasc Imaging* 2007;23(3):361-7. *Full-text exclusion reason(s): No outcomes of interest.*

Becker A, Leber AW, Becker C, et al. Predictive value of coronary calcifications for future cardiac events in asymptomatic patients with diabetes mellitus: a prospective study in 716 patients over 8 years. *BMC Cardiovasc Disord* 2008;8:27. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Study does not compare one NIT to another or to diagnostic cardiac catheterization.*

Becker CR, Knez A, Leber A, et al. Detection of coronary artery stenoses with multislice helical CT angiography. *J Comput Assist Tomogr* 2002;26(5):750-5. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup; No outcomes of interest.*

Bedetti G, Gargani L, Sicari R, et al. Comparison of prognostic value of echographic [corrected] risk score with the Thrombolysis in Myocardial Infarction (TIMI) and Global Registry in Acute Coronary Events (GRACE) risk scores in acute coronary syndrome. *Am J Cardiol* 2010;106(12):1709-16. *Full-text exclusion reason(s): No data for NITs of interest (ECG, ECHO, SPECT, PET, CMR, CTA)*

Bedetti G, Pasanisi EM, Pizzi C, et al. Economic analysis including long-term risks and costs of alternative diagnostic strategies to evaluate patients with chest pain. *Cardiovasc Ultrasound* 2008;6:21. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry; Data for women not reported as a subgroup; No outcomes of interest.*

Bedetti G, Pasanisi EM, Tintori G, et al. Stress echo in chest pain unit: the SPEED trial. *Int J Cardiol* 2005;102(3):461-7. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Beer SG, Heo J, Kong B, et al. Use of oral dipyridamole SPECT thallium-201 imaging in detection of coronary artery disease. *Am Heart J* 1989;118(5 Pt 1):1022-7. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Beigel R, Oieru D, Goitein O, et al. Usefulness of routine use of multidetector coronary computed tomography in the "fast track" evaluation of patients with acute chest pain. *Am J Cardiol* 2009;103(11):1481-6. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Beinart R, Matetzky S, Shechter M, et al. Stress-induced ST-segment elevation in patients without prior Q-wave myocardial infarction. *J Electrocardiol* 2008;41(4):312-7. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Beleslin BD, Ostojic M, Djordjevic-Dikic A, et al. Integrated evaluation of relation between coronary lesion features and stress echocardiography results: the importance of coronary lesion morphology. *J Am Coll Cardiol* 1999;33(3):717-26. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Beleslin BD, Ostojic M, Stepanovic J, et al. Stress echocardiography in the detection of myocardial ischemia. Head-to-head comparison of exercise, dobutamine, and dipyridamole tests. *Circulation* 1994;90(3):1168-76. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Belge B, Coche E, Pasquet A, et al. Accurate estimation of global and regional cardiac function by retrospectively gated multidetector row computed tomography: comparison with cine magnetic resonance imaging. *Eur Radiol* 2006;16(7):1424-33. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Bell GW, Edwardes M, Dunning AM, et al. Periprocedural safety of 64-detector row coronary computed tomographic angiography: results from the prospective multicenter ACCURACY trial. *J Cardiovasc Comput Tomogr* 2010;4(6):375-80. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Bellasi A, Ferramosca E, Muntner P, et al. Correlation of simple imaging tests and coronary artery calcium measured by computed tomography in hemodialysis patients. *Kidney Int* 2006;70(9):1623-8. *Full-text exclusion*

*reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup; No outcomes of interest.*

Beller GA. Perfusion imaging. *J Am Coll Cardiol* 2000;35(5 SUPPL. B):29B-31B. *Full-text exclusion reason(s): Not a clinical study report.*

Ben-Gal T, Zafirir N. The utility and potential cost-effectiveness of stress myocardial perfusion thallium SPECT imaging in hospitalized patients with chest pain and normal or non-diagnostic electrocardiogram. *Isr Med Assoc J* 2001;3(10):725-30. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Ben-Gal T, Zafirir N. Utility of stress myocardial perfusion imaging in hospitalized patients with chest pain and normal or nondiagnostic electrocardiograms. *Cardiovascular Reviews and Reports* 2001;22(10):600-606. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Berger BC, Watson DD, Taylor GJ, et al. Quantitative thallium-201 exercise scintigraphy for detection of coronary artery disease. *J Nucl Med* 1981;22(7):585-93. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Bergeron S, Ommen SR, Bailey KR, et al. Exercise echocardiographic findings and outcome of patients referred for evaluation of dyspnea. *J Am Coll Cardiol* 2004;43(12):2242-6. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Berman D, Hachamovitch R, Lewin H, et al. Risk stratification in coronary artery disease: Implications for stabilization and prevention. *Am J Cardiol* 1997;79(12 B):10-16. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Berman DS, Hachamovitch R, Shaw LJ, et al. Roles of nuclear cardiology, cardiac computed tomography, and cardiac magnetic resonance: Noninvasive risk stratification and a conceptual framework for the selection of noninvasive imaging tests in patients with known or suspected coronary artery disease. *J Nucl Med* 2006;47(7):1107-18. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Berman DS, Kang X, Gransar H, et al. Quantitative assessment of myocardial perfusion abnormality on SPECT myocardial perfusion imaging is more reproducible than expert visual analysis. *J Nucl Cardiol* 2009;16(1):45-53. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Berman DS, Kang X, Hayes SW, et al. Adenosine myocardial perfusion single-photon emission computed tomography in women compared with men. Impact of diabetes mellitus on incremental prognostic value and effect on patient management. *J Am Coll Cardiol* 2003;41(7):1125-33. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization.*

Berman DS, Kang X, Schisterman EF, et al. Serial changes on quantitative myocardial perfusion SPECT in patients undergoing revascularization or conservative therapy. *J Nucl Cardiol* 2001;8(4):428-37. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup; No outcomes of interest.*

Berman DS, Kiat H, Friedman JD, et al. Separate acquisition rest thallium-201/stress technetium-99m sestamibi dual-isotope myocardial perfusion single-photon emission computed tomography: a clinical validation study. *J Am Coll Cardiol* 1993;22(5):1455-64. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Berman DS, Wong ND, Gransar H, et al. Relationship between stress-induced myocardial ischemia and atherosclerosis measured by coronary calcium tomography. *J Am Coll Cardiol* 2004;44(4):923-30. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Bernhardt P, Engels T, Levenson B, et al. Prediction of necessity for coronary artery revascularization by adenosine contrast-enhanced magnetic resonance imaging. *Int J Cardiol* 2006;112(2):184-90. *Full-text exclusion reason(s): No*

women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.

Bernhardt P, Levenson B, Albrecht A,, et al. Detection of cardiac small vessel disease by adenosine-stress magnetic resonance. *Int J Cardiol* 2007;121(3):261-6. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Bernhardt P, Manzke R, Bornstedt A,, et al. Blood oxygen level-dependent magnetic resonance imaging using T2-prepared steady-state free-precession imaging in comparison to contrast-enhanced myocardial perfusion imaging. *Int J Cardiol* 2011;147(3):416-419. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Bernheim AM, Kittipovanonth M, Scott CG,, et al. Relation of dyspnea in patients unable to perform exercise stress testing to outcome and myocardial ischemia. *Am J Cardiol* 2009;104(2):265-9. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Bernheim AM, Kittipovanonth M, Takahashi PY,, et al. Does the prognostic value of dobutamine stress echocardiography differ among different age groups? *Am Heart J* 2011;161(4):740-5. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Beslic N, Kucukalic-Selimovic E. Comparison of the diagnostic capabilities of noninvasive methods for early detection of coronary artery disease. *Med Arh* 2011;65(2):96-8. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Bestetti A, Di Leo C, Alessi A,, et al. Post-stress end-systolic left ventricular dilation: a marker of endocardial post-ischemic stunning. *Nucl Med Commun* 2001;22(6):685-93. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Bettencourt N, Rocha J, Carvalho M,, et al. Multislice computed tomography in the exclusion of coronary artery disease in patients with presurgical valve disease. *Circ Cardiovasc Imaging* 2009;2(4):306-13. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Bhat A, Desai A, Amsterdam EA. Usefulness of high functional capacity in patients with exercise-induced ST-depression to predict a negative result on exercise echocardiography and low prognostic risk. *Am J Cardiol* 2008;101(11):1541-3. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Bholasingh R, Cornel JH, Kamp O,, et al. Prognostic value of predischARGE dobutamine stress echocardiography in chest pain patients with a negative cardiac troponin T. *J Am Coll Cardiol* 2003;41(4):596-602. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Biagini E, Elhendy A, Bax JJ,, et al. Seven-year follow-up after dobutamine stress echocardiography: impact of gender on prognosis. *J Am Coll Cardiol* 2005;45(1):93-7. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization.*

Bielak LF, Rumberger JA, Sheedy PF, 2nd,, et al. Probabilistic model for prediction of angiographically defined obstructive coronary artery disease using electron beam computed tomography calcium score strata. *Circulation* 2000;102(4):380-5. *Full-text exclusion reason(s): No data for NITs of interest (ECG, ECHO, SPECT, PET, CMR, CTA).*

Bigi R, Bax JJ, van Domburg RT,, et al. Simultaneous echocardiography and myocardial perfusion single photon emission computed tomography associated with dobutamine stress to predict long-term cardiac mortality in normotensive and hypertensive patients. *J Hypertens* 2005;23(7):1409-15. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Bigi R, Desideri A, Cortigiani L,, et al. Stress echocardiography for risk stratification of diabetic patients with known or suspected coronary artery disease. *Diabetes Care* 2001;24(9):1596-601. *Full-text exclusion reason(s):*



*Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Bigi R, Occhi G, Fiorentini C, et al. Dobutamine stress echocardiography for the identification of multivessel coronary artery disease after uncomplicated myocardial infarction: the importance of test end-point. *Int J Cardiol* 1995;50(1):51-60. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup*

Bingham SE, Hachamovitch R. Incremental prognostic significance of combined cardiac magnetic resonance imaging, adenosine stress perfusion, delayed enhancement, and left ventricular function over preimaging information for the prediction of adverse events. *Circulation* 2011;123(14):1509-18. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Bischoff B, Hein F, Meyer T, et al. Comparison of sequential and helical scanning for radiation dose and image quality: results of the Prospective Multicenter Study on Radiation Dose Estimates of Cardiac CT Angiography (PROTECTION) I Study. *AJR Am J Roentgenol*; 2010:1495-9. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Bjornstad K, Aakhus S, Hatle L. Digital high frame rate stress echocardiography for detection of coronary artery stenosis by high dose dipyridamole stress testing. *Int J Card Imaging* 1995;11(3):163-70. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Bjornstad K, Aakhus S, Torp HG. How does computer-assisted digital wall motion analysis influence observer agreement and diagnostic accuracy during stress echocardiography? *Int J Card Imaging* 1997;13(2):105-14. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Blankstein R, Devore AD. Selecting a noninvasive imaging study after an inconclusive exercise test. *Circulation* 2010;122(15):1514-1518. *Full-text exclusion reason(s): Not a clinical study report.*

Blankstein R, Di Carli MF. Integration of coronary anatomy and myocardial perfusion imaging. *Nature Reviews Cardiology* 2010;7(4):226-236. *Full-text exclusion reason(s): Not a clinical study report.*

Blankstein R, Okada DR, Rocha-Filho JA, et al. Cardiac myocardial perfusion imaging using dual source computed tomography. *Int J Cardiovasc Imaging* 2009;25(SUPPL. 2):209-216. *Full-text exclusion reason(s): Not a clinical study report.*

Blankstein R, Shturman LD, Rogers IS, et al. Adenosine-induced stress myocardial perfusion imaging using dual-source cardiac computed tomography. *J Am Coll Cardiol* 2009;54(12):1072-84. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Blinder G, Benhorin J, Koukoui D, et al. The value of electrocardiography-gated multi-slice computed tomography in the evaluation of patients with chest pain. *Isr Med Assoc J* 2005;7(7):419-23. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Blomkalns AL, Chen AY, Hochman JS, et al. Gender disparities in the diagnosis and treatment of non-ST-segment elevation acute coronary syndromes: large-scale observations from the CRUSADE (Can Rapid Risk Stratification of Unstable Angina Patients Suppress Adverse Outcomes With Early Implementation of the American College of Cardiology/American Heart Association Guidelines) National Quality Improvement Initiative. *J Am Coll Cardiol* 2005;45(6):832-7. *Full-text exclusion reason(s): No data for NITs of interest (ECG, ECHO, SPECT, PET, CMR, CTA).*

Blumenthal RS, Becker DM, Yanek LR, et al. Detecting occult coronary disease in a high-risk asymptomatic population. *Circulation* 2003;107(5):702-7. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Blumenthal RS, Becker DM, Yanek LR, et al. Comparison of coronary calcium and stress myocardial perfusion imaging in apparently healthy siblings of individuals with premature coronary artery disease. *Am J Cardiol* 2006;97(3):328-33. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Bodenheimer MM, Banka VS, Fooshee CM, et al. Comparative sensitivity of the exercise electrocardiogram, thallium imaging and stress radionuclide angiography to detect the presence and severity of coronary heart disease.

Circulation 1979;60(6):1270-8. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Bodi V, Sanchis J, Lopez-Lereu MP, et al. Prognostic value of dipyridamole stress cardiovascular magnetic resonance imaging in patients with known or suspected coronary artery disease. J Am Coll Cardiol 2007;50(12):1174-9. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Bodi V, Sanchis J, Lopez-Lereu MP, et al. Prognostic and therapeutic implications of dipyridamole stress cardiovascular magnetic resonance on the basis of the ischaemic cascade. Heart 2009;95(1):49-55. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Bogaert J, Kuzo R, Dymarkowski S, et al. Coronary artery imaging with real-time navigator three-dimensional turbo-field-echo MR coronary angiography: initial experience. Radiology 2003;226(3):707-16. *Full-text exclusion reason(s): All women in the study are known to have CAD; No outcomes of interest.*

Bokhari S, Blood DK, Bergmann SR. Failure of right precordial electrocardiography during stress testing to identify coronary artery disease. J Nucl Cardiol 2001;8(3):325-31. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Bonmassari R, Muraglia S, Centonze M, et al. Noninvasive detection of coronary artery stenosis with 16-slice spiral computed tomography in a population at low to moderate risk for coronary artery disease. J Cardiovasc Med (Hagerstown) 2006;7(11):817-25. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Boogers MJ, Schuijf JD, Kitslaar PH, et al. Automated quantification of stenosis severity on 64-slice CT: A comparison with quantitative coronary angiography. JACC: Cardiovascular Imaging 2010;3(7):699-709. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Bordeleau E, Lamonde A, Prenovault J, et al. Accuracy and rate of coronary artery segment visualization with CT angiography for the non-invasive detection of coronary artery stenoses. Int J Cardiovasc Imaging 2007;23(6):771-80. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Borer JS, Brensike JF, Redwood DR, et al. Limitations of the electrocardiographic response to exercise in predicting coronary-artery disease. N Engl J Med 1975;293(8):367-71. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Borges-Neto S, Shaw LK, Tuttle RH, et al. Incremental prognostic power of single-photon emission computed tomographic myocardial perfusion imaging in patients with known or suspected coronary artery disease. Am J Cardiol 2005;95(2):182-8. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Borges-Neto S, Tuttle RH, Shaw LK, et al. Outcome prediction in patients at high risk for coronary artery disease: comparison between 99mTc tetrofosmin and 99mTc sestamibi. Radiology 2004;232(1):58-65. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Boshchenko AA, Vrublevsky AV, Karpov RS. Transthoracic echocardiography in the detection of chronic total coronary artery occlusion. Eur J Echocardiogr 2009;10(1):62-8. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup; No outcomes of interest.*

Boudik F, Anger Z, Aschermann M, et al. Dipyridamole body surface potential mapping: noninvasive differentiation of syndrome X from coronary artery disease. J Electrocardiol 2002;35(3):181-91. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup; No outcomes of interest.*

Boudoulas H, Weissler AM, Sohn YH, et al. Detection of the High Mortality Risk by Combined Noninvasive Measures of Mechanical and Electrical Dysfunction in Coronary Artery Disease. Joint Meeting of the American Federation for Clinical Research, the Association of American Physicians and the American Society for Clinical Investigation, Washington, DC, 7-10 May 82 (World Meeting Number 822 0218). *Full-text exclusion reason(s): Full-text unobtainable.*

Bourque JM, Charlton GT, Holland BH, et al. Prognosis in patients achieving  $\geq 10$  METS on exercise stress testing: was SPECT imaging useful? *J Nucl Cardiol* 2011;18(2):230-7. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Bourque JM, Holland BH, Watson DD, et al. Achieving an exercise workload of  $>$  or  $= 10$  metabolic equivalents predicts a very low risk of inducible ischemia: does myocardial perfusion imaging have a role? *J Am Coll Cardiol* 2009;54(6):538-45. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup; No outcomes of interest.*

Boussel L, Gamondes D, Staat P, et al. Acute chest pain with normal coronary angiogram: role of contrast-enhanced multidetector computed tomography in the differential diagnosis between myocarditis and myocardial infarction. *J Comput Assist Tomogr* 2008;32(2):228-32. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Bouvier E, Logeart D, Sablayrolles JL, et al. Diagnosis of aortic valvular stenosis by multislice cardiac computed tomography. *Euro Heart J* 2006(24):3033-8. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup; No outcomes of interest.*

Bouzas-Mosquera A, Peteiro J, Alvarez-Garcia N, et al. Prediction of mortality and major cardiac events by exercise echocardiography in patients with normal exercise electrocardiographic testing. *J Am Coll Cardiol* 2009;53(21):1981-90. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Boyne TS, Koplan BA, Parsons WJ, et al. Predicting adverse outcome with exercise SPECT technetium-99m sestamibi imaging in patients with suspected or known coronary artery disease. *Am J Cardiol* 1997;79(3):270-4. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Brodofel H, Burgstahler C, Tsiflikas I, et al. Dual-source CT: effect of heart rate, heart rate variability, and calcification on image quality and diagnostic accuracy. *Radiology* 2008;247(2):346-55. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Brook RD, Bard RL, Patel S, et al. A negative carotid plaque area test is superior to other noninvasive atherosclerosis studies for reducing the likelihood of having underlying significant coronary artery disease. *Arterioscler Thromb Vasc Biol* 2006;26(3):656-62. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Brown KA. Prognostic value of thallium-201 myocardial perfusion imaging. A diagnostic tool comes of age. *Circulation* 1991;83(2):363-81. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Brown KA, Boucher CA, Okada RD, et al. Prognostic value of exercise thallium-201 imaging in patients presenting for evaluation of chest pain. *J Am Coll Cardiol* 1983;1(4):994-1001. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Brown RA, Schlegel TT. Diagnostic utility of the spatial versus individual planar QRS-T angles in cardiac disease detection. *J Electrocardiol* 2011;44(4):404-409. *Full-text exclusion reason(s): No data for NITs of interest (ECG, ECHO, SPECT, PET, CMR, CTA).*

Brown TL, Voicu C, Merrill J, et al. Pathophysiologic correlates of  $^{82}\text{Rb}$  biodistribution in cardiac PET/CT. *Eur J Nucl Med Mol Imaging* 2011;38(3):479-84. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup; No outcomes of interest.*

Bruder O, Schneider S, Nothnagel D, et al. EuroCMR (European Cardiovascular Magnetic Resonance) registry: results of the German pilot phase. *J Am Coll Cardiol* 2009;54(15):1457-66. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Bucci M, Joutsiniemi E, Saraste A, et al. Intrapericardial, but not extrapericardial, fat is an independent predictor of impaired hyperemic coronary perfusion in coronary artery disease. *Arterioscler Thromb Vasc Biol* 2011;31(1):211-8. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Bucerius J, Joe AY, Herder E, et al. Hemodynamic variables during stress testing can predict referral to early catheterization but failed to show a prognostic impact on emerging cardiac events in patients aged 70 years and older undergoing exercise (99m)Tc-sestamibi myocardial perfusion scintigraphy. *Int J Cardiovasc Imaging* 2009;25(6):569-79. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Bucerius J, Joe AY, Herder E, et al. Pathological 99mTc-sestamibi myocardial perfusion scintigraphy is independently associated with emerging cardiac events in elderly patients with known or suspected coronary artery disease. *Acta Radiol* 2011;52(1):52-8. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Bucerius J, Joe AY, Herder E, et al. Significant association of female gender with lower degree of pathological 99mTc-sestamibi scintigraphy results as well as higher cardiac-related deaths free survival in elderly patients. *Med Klin (Munich)* 2010;105(12):901-9. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization.*

Buchthal SD, den Hollander JA, Merz CN, et al. Abnormal myocardial phosphorus-31 nuclear magnetic resonance spectroscopy in women with chest pain but normal coronary angiograms. *N Engl J Med* 2000;342(12):829-35. *Full-text exclusion reason(s): All women in the study are known to have CAD; No outcomes of interest.*

Budoff MJ. Coronary CT Angiography after invasive angiography: is it worth it? *J Invasive Cardiol* 2008;20(1):7-8. *Full-text exclusion reason(s): Not a clinical study report.*

Budoff MJ, Ahmadi N, Sarraf G, et al. Determination of left ventricular mass on cardiac computed tomographic angiography. *Acad Radiol* 2009;16(6):726-32. *Full-text exclusion reason(s): No outcomes of interest.*

Budoff MJ, Diamond GA, Raggi P, et al. Continuous probabilistic prediction of angiographically significant coronary artery disease using electron beam tomography. *Circulation* 2002;105(15):1791-6. *Full-text exclusion reason(s): No data for NITs of interest (ECG, ECHO, SPECT, PET, CMR, CTA).*

Budoff MJ, Dowe D, Jollis JG, et al. Diagnostic performance of 64-multidetector row coronary computed tomographic angiography for evaluation of coronary artery stenosis in individuals without known coronary artery disease: results from the prospective multicenter ACCURACY (Assessment by Coronary Computed Tomographic Angiography of Individuals Undergoing Invasive Coronary Angiography) trial. *J Am Coll Cardiol* 2008;52(21):1724-32. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Budoff MJ, Georgiou D, Brody A, et al. Ultrafast computed tomography as a diagnostic modality in the detection of coronary artery disease: a multicenter study. *Circulation* 1996;93(5):898-904. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Budoff MJ, Gopal A, Gopalakrishnan D. Cardiac computed tomography: Diagnostic utility and integration in clinical practice. *Clin Cardiol* 2006;29(9 SUPPL. 1):I4-I14. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Budoff MJ, Gopal A, Gul KM,, et al. Prevalence of obstructive coronary artery disease in an outpatient cardiac CT angiography environment. *Int J Cardiol* 2008;129(1):32-6. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Budoff MJ, Lu B, Shinbane JS,, et al. Methodology for improved detection of coronary stenoses with computed tomographic angiography. *Am Heart J* 2004;148(6):1085-90. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Budoff MJ, Rasouli ML, Shavelle DM,, et al. Cardiac CT angiography (CTA) and nuclear myocardial perfusion imaging (MPI)-a comparison in detecting significant coronary artery disease. *Acad Radiol* 2007;14(3):252-7. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Budoff MJ, Shavelle DM, Lamont DH,, et al. Usefulness of electron beam computed tomography scanning for distinguishing ischemic from nonischemic cardiomyopathy. *J Am Coll Cardiol* 1998;32(5):1173-8. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD.*

Budoff MJ, Shokooh S, Shavelle RM,, et al. Electron beam tomography and angiography: sex differences. *Am Heart J* 2002;143(5):877-82. *Full-text exclusion reason(s): No data for NITs of interest (ECG, ECHO, SPECT, PET, CMR, CTA).*

Budzynski J. Does esophageal dysfunction affect the course of treadmill stress test in patients with recurrent angina-like chest pain? *Pol Arch Med Wewn* 2010;120(12):484-90. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup; No outcomes of interest.*

Buechel RR, Pazhenkottil AP, Herzog BA,, et al. Prognostic performance of low-dose coronary CT angiography with prospective ECG triggering. *Heart* 2011;97(17):1385-1390. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization.*

Bunce NH, Jhooti P, Keegan J,, et al. Evaluation of free-breathing three-dimensional magnetic resonance coronary angiography with hybrid ordered phase encoding (HOPE) for the detection of proximal coronary artery stenosis. *J Magn Reson Imaging* 2001;14(6):677-84. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Bunce NH, Reyes E, Keegan J,, et al. Combined coronary and perfusion cardiovascular magnetic resonance for the assessment of coronary artery stenosis. *J Cardiovasc Magn Reson* 2004;6(2):527-39. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Bungo MW, Leland OS, Jr. Discordance of exercise thallium testing with coronary arteriography in patients with atypical presentations. *Chest* 1983;83(1):112-6. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Burgstahler C, Beck T, Kuettner A,, et al. Image quality and diagnostic accuracy of 16-slice multidetector computed tomography for the detection of coronary artery disease in obese patients. *Int J Obes (Lond)* 2006;30(3):569-73. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Burgstahler C, Beck T, Kuettner A,, et al. Image quality and diagnostic accuracy of 16-slice multidetector spiral computed tomography for the detection of coronary artery disease in elderly patients. *J Comput Assist Tomogr* 2005;29(6):734-8. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Burgstahler C, Beck T, Reimann A,, et al. Diagnostic accuracy of multislice computed tomography for the detection of coronary artery disease in diabetic patients. *J Diabetes Complications* 2007;21(2):69-74. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Burgstahler C, Kunze M, Gawaz MP,, et al. Adenosine stress first pass perfusion for the detection of coronary artery disease in patients with aortic stenosis: a feasibility study. *Int J Cardiovasc Imaging* 2008;24(2):195-200. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Burgstahler C, Reimann A, Drosch T., et al. Cardiac dual-source computed tomography in patients with severe coronary calcifications and a high prevalence of coronary artery disease. *J Cardiovasc Comput Tomogr* 2007;1(3):143-51. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Burns RJ, Galligan L, Wright LM., et al. Improved specificity of myocardial thallium-201 single-photon emission computed tomography in patients with left bundle branch block by dipyridamole. *Am J Cardiol* 1991;68(5):504-8. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Butler J, Shapiro M, Reiber J., et al. Extent and distribution of coronary artery disease: a comparative study of invasive versus noninvasive angiography with computed angiography. *Am Heart J* 2007;153(3):378-84. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Bybee KA, Lee J, Markiewicz R., et al. Diagnostic and clinical benefit of combined coronary calcium and perfusion assessment in patients undergoing PET/CT myocardial perfusion stress imaging. *J Nucl Cardiol* 2010;17(2):188-196. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Cademartiri F, Carli MF. MSCT is better than stress perfusion imaging for detecting CAD. *European Journal of Nuclear Medicine and Molecular Imaging* 2006;33(3):353-359. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Cademartiri F, La Grutta L, Malago R., et al. Assessment of left main coronary artery atherosclerotic burden using 64-slice CT coronary angiography: correlation between dimensions and presence of plaques. *Radiol Med* 2009;114(3):358-69. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Cademartiri F, La Grutta L, Palumbo A., et al. Computed tomography coronary angiography vs. stress ECG in patients with stable angina. *Radiol Med* 2009;114(4):513-23. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Cademartiri F, Maffei E, Notarangelo F., et al. 64-slice computed tomography coronary angiography: diagnostic accuracy in the real world. *Radiol Med* 2008;113(2):163-80. *Full-text exclusion reason(s): All women in the study are known to have CAD; No outcomes of interest.*

Cademartiri F, Maffei E, Palumbo A., et al. Diagnostic accuracy of 64-slice computed tomography coronary angiography in patients with low-to-intermediate risk. *Radiol Med* 2007;112(7):969-81. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Cademartiri F, Maffei E, Palumbo A., et al. Diagnostic accuracy of computed tomography coronary angiography in patients with a zero calcium score. *Eur Radiol* 2010;20(1):81-7. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Cademartiri F, Maffei E, Palumbo A., et al. Coronary calcium score and computed tomography coronary angiography in high-risk asymptomatic subjects: assessment of diagnostic accuracy and prevalence of non-obstructive coronary artery disease. *Eur Radiol* 2010;20(4):846-54. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Cademartiri F, Maffei E, Palumbo AA., et al. Influence of intra-coronary enhancement on diagnostic accuracy with 64-slice CT coronary angiography. *Eur Radiol* 2008;18(3):576-83. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Cademartiri F, Malago R, Belgrano M., et al. Spectrum of collateral findings in multislice CT coronary angiography. *Radiol Med* 2007;112(7):937-48. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup; No outcomes of interest.*

Cademartiri F, Mollet NR, Lemos PA., et al. Impact of coronary calcium score on diagnostic accuracy for the detection of significant coronary stenosis with multislice computed tomography angiography. *Am J Cardiol* 2005;95(10):1225-7. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Cademartiri F, Mollet NR, Runza G,, et al. Diagnostic accuracy of multislice computed tomography coronary angiography is improved at low heart rates. *Int J Cardiovasc Imaging* 2006;22(1):101-5; discussion 107-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Cademartiri F, Romano M, Seitun S,, et al. Prevalence and characteristics of coronary artery disease in a population with suspected ischemic heart disease using CT coronary angiography: correlations with cardiovascular risk factors and clinical presentation. *Radiol Med* 2008;113(3):363-72. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Cademartiri F, Runza G, Marano R,, et al. Diagnostic accuracy of 16-row multislice CT angiography in the evaluation of coronary segments. *Radiol Med* 2005;109(1-2):91-7. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Cademartiri F, Seitun S, Romano M,, et al. Prognostic value of 64-slice coronary angiography in diabetes mellitus patients with known or suspected coronary artery disease compared with a nondiabetic population. *Radiol Med* 2008;113(5):627-43. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Campisi R. Noninvasive assessment of coronary microvascular function in women at risk for ischaemic heart disease. *Int J Clin Pract* 2008;62(2):300-7. *Full-text exclusion reason(s): Not a clinical study report.*

Campos AM, da Cunha AB. Dobutamine stress echocardiography as a predictor of coronary lesion severity on coronary angiography. *Rev Port Cardiol* 2007;26(5):505-18. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Candell-Riera J, Oller-Martinez G, de Leon G,, et al. Yield of early rest and stress myocardial perfusion single-photon emission computed tomography and electrocardiographic exercise test in patients with atypical chest pain, nondiagnostic electrocardiogram, and negative biochemical markers in the emergency department. *Am J Cardiol* 2007;99(12):1662-6. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Candell-Riera J, Oller-Martínez G, Pereztol-Valdés O,, et al. Early myocardial perfusion gated-SPECT in patients with chest pain and non-diagnostic ECG in the emergency department. *Revista española de cardiología* 2004(3):225-33. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Candell-Riera J, Santana-Boado C, Bermejo B,, et al. Interhospital observer agreement in interpretation of exercise myocardial Tc-99m tetrofosmin SPECT studies. *J Nucl Cardiol* 2001;8(1):49-57. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Carmo MM, Ferreira T, Quininha J,, et al. Non-invasive coronary artery evaluation with multidetector computed tomography. *Rev Port Cardiol* 2005;24(5):667-79. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Carrascosa P, Capunay C, Bettinotti M,, et al. Feasibility of gadolinium-diethylene triamine pentaacetic acid enhanced multidetector computed tomography for the evaluation of coronary artery disease. *J Cardiovasc Comput Tomogr* 2007;1(2):86-94. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Carrascosa P, Capunay C, Deviggiano A,, et al. Feasibility of 64-slice gadolinium-enhanced cardiac CT for the evaluation of obstructive coronary artery disease. *Heart* 2010;96(19):1543-1549. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Carrascosa P, Capunay C, Deviggiano A,, et al. Accuracy of low-dose prospectively gated axial coronary CT angiography for the assessment of coronary artery stenosis in patients with stable heart rate. *J Cardiovasc Comput Tomogr* 2010;4(3):197-205. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Carrascosa PM, Capunay CM, Parodi JC,, et al. General utilities of multislice tomography in the cardiac field. *Herz* 2003;28(1):44-51. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Carrigan TP, Nair D, Schoenhagen P, et al. Prognostic utility of 64-slice computed tomography in patients with suspected but no documented coronary artery disease. *Eur Heart J* 2009;30(3):362-71. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Carrillof Lopez L, Frade Garcia J, Gomez Lepe A, et al. [Correlation between findings of exertion ECG with 12 derivations and coronariography]. *Arch Inst Cardiol Mex* 1978;48(4):823-39. *Full-text exclusion reason(s): Non-English.*

Carrinho M, Moraes A, Morcerf F, et al. Myocardial contrast echocardiography in patients with suspected or known coronary artery disease: comparison with myocardial nuclear scintigraphy. *Arq Bras Cardiol* 2004;83(5):419-23; 414-8. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Caruana L, Petrie MC, Davie AP, et al. Do patients with suspected heart failure and preserved left ventricular systolic function suffer from "diastolic heart failure" or from misdiagnosis? A prospective descriptive study. *BMJ* 2000;321(7255):215-8. *Full-text exclusion reason(s): No data for NITs of interest (ECG, ECHO, SPECT, PET, CMR, CTA).*

Casolo G, Del Meglio J, Rega L, et al. Detection and assessment of coronary artery anomalies by three-dimensional magnetic resonance coronary angiography. *Int J Cardiol* 2005;103(3):317-22. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup; No outcomes of interest.*

Castini D, Gentile F, Ornaghi M, et al. Dobutamine echocardiography: usefulness of digital image processing. *Eur Heart J* 1995;16(10):1420-4. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Catalan P, Leta R, Hidalgo A, et al. Ruling out coronary artery disease with noninvasive coronary multidetector CT angiography before noncoronary cardiovascular surgery. *Radiology* 2011;258(2):426-34. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Caussin C, Daoud B, Ghostine S, et al. Comparison of lumens of intermediate coronary stenosis using 16-slice computed tomography versus intravascular ultrasound. *Am J Cardiol* 2005;96(4):524-8. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Caussin C, Ohanessian A, Ghostine S, et al. Characterization of vulnerable nonstenotic plaque with 16-slice computed tomography compared with intravascular ultrasound. *Am J Cardiol* 2004;94(1):99-104. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization.*

Cecil MP, Kosinski AS, Jones MT, et al. The importance of work-up (verification) bias correction in assessing the accuracy of SPECT thallium-201 testing for the diagnosis of coronary artery disease. *J Clin Epidemiol* 1996;49(7):735-42. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Cerci MSJ, Cerci JJ, Cerci RJ, et al. Myocardial perfusion imaging is a strong predictor of death in women. *JACC: Cardiovascular Imaging* 2011;4(8):880-888. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization.*

Chalela WA, Mansur AP, Aldrighi JM. Noninvasive diagnostic evaluation for chest pain in women. *Arq Bras Cardiol* 2001;76(6):535-44. *Full-text exclusion reason(s): Not a clinical study report.*

Chammas E, Yatim A, Hage C, et al. Evaluation of Tc-99m tetrofosmin scan for coronary artery disease diagnosis. *Asian Cardiovasc Thorac Ann* 2002;10(3):244-7. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Chan RK, Tonkin AM, Byrgiotis S, et al. Assessment of coronary artery disease by dobutamine stress echocardiography (DSE). *Aust N Z J Med* 1995;25(6):707-15. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Chang AM, Shofer FS, Weiner MG, et al. Actual financial comparison of four strategies to evaluate patients with potential acute coronary syndromes. *Acad Emerg Med* 2008;15(7):649-55. *Full-text exclusion reason(s): No women*



*with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup; No outcomes of interest.*

Chang SA, Choi SI, Choi EK, et al. Usefulness of 64-slice multidetector computed tomography as an initial diagnostic approach in patients with acute chest pain. *Am Heart J* 2008;156(2):375-83. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup*

Chang SM, Hakeem A, Nagueh SF. Predicting clinically unrecognized coronary artery disease: use of two-dimensional echocardiography. *Cardiovasc Ultrasound* 2009;7:10. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Chang SM, Nabi F, Xu J, et al. The coronary artery calcium score and stress myocardial perfusion imaging provide independent and complementary prediction of cardiac risk. *J Am Coll Cardiol* 2009;54(20):1872-82. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Chang SM, Nabi F, Xu J, et al. Normal stress-only versus standard stress/rest myocardial perfusion imaging: similar patient mortality with reduced radiation exposure. *J Am Coll Cardiol* 2010;55(3):221-30. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup; No outcomes of interest.*

Chao SP, Law WY, Kuo CJ, et al. The diagnostic accuracy of 256-row computed tomographic angiography compared with invasive coronary angiography in patients with suspected coronary artery disease. *Euro Heart J* 2010;31(15):1916-1923. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Charvat J, Michalova K, Taborska K, et al. Comparison of the exercise ECG and stress myocardial SPECT in detection of the significant coronary artery disease in the asymptomatic patients with diabetes mellitus type 2. *Bratisl Lek Listy* 2004;105(2):56-61. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Chatranukulchai P, Tumkosit M, Cholteesupachai J, et al. Diagnostic accuracy of combined dipyridamole stress perfusion and delayed enhancement cardiovascular magnetic resonance imaging for detection of coronary artery disease. *Asian Biomedicine* 2010;4(1):19-25. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Chatziioannou SN, Moore WH, Dhekne RD, et al. Women with high exercise tolerance and the role of myocardial perfusion imaging. *Clin Cardiol* 2001;24(6):475-80. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup.*

Chatziioannou SN, Moore WH, Ford PV, et al. Prognostic value of myocardial perfusion imaging in patients with high exercise tolerance. *Circulation* 1999;99(7):867-72. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Chaudhry FA, Tauke JT, Alessandrini RS, et al. Enhanced detection of ischemic myocardium by transesophageal dobutamine stress echocardiography: comparison with simultaneous transthoracic echocardiography. *Echocardiography* 2000;17(3):241-53. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Cheezum MK, Hulten EA, Taylor AJ, et al. Cardiac CT angiography compared with myocardial perfusion stress testing on downstream resource utilization. *J Cardiovasc Comput Tomogr* 2011;5(2):101-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Chen G, Redberg RF. Noninvasive diagnostic testing of coronary artery disease in women. *Cardiol Rev* 2000;8(6):354-60. *Full-text exclusion reason(s): Not a clinical study report.*

Chen HW, Fang XM, Hu XY, et al. Efficacy of dual-source CT coronary angiography in evaluating coronary stenosis: initial experience. *Clin Imaging* 2010;34(3):165-71. *Full-text exclusion reason(s): No women with*

*symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup; No outcomes of interest.*

Chen JC, Wang MT, Wang YIC, et al. 40-Slice multi-detector computed tomography in detecting coronary artery stenosis: A preliminary result. *Chinese Journal of Radiology* 2009;34(3):161-170+171. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Chen LC, Chen JW, Wu MH, et al. Differential coronary artery calcification detected by electron beam computed tomography as an indicator of coronary stenosis among patients with stable angina pectoris. *Can J Cardiol* 2001;17(6):667-76. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Chen LC, Ding PY, Chen JW, et al. Coronary artery calcium determined by electron beam computed tomography for predicting angiographic coronary artery disease in moderate- to high-risk Chinese patients. *Cardiology* 2001;95(4):183-9. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Chen Y, Han P, Liang B, et al. Comparative study on 16-slice CT coronary angiography vs conventional coronary angiography—a report of 38 cases. *J Huazhong Univ Sci Technolog Med Sci* 2008;28(1):110-3. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Chen YC, Chin KC, Ko JT. Value of ECG-gated thallium-201 dipyridamole SPECT in borderline cases of myocardial perfusion scan. *Acta Cardiologica Sinica* 2006;22(1):24-30. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Chen Z, Duan Q, Xue X, et al. Noninvasive detection of coronary artery stenoses with contrast-enhanced whole-heart coronary magnetic resonance angiography at 3.0 T. *Cardiology* 2010;117(4):284-290. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Cheneau E, Vahdat B, Bernard L, et al. Routine use of coronary computed tomography as initial diagnostic test for angina pectoris. *Arch Cardiovasc Dis* 2011;104(1):29-34. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Cheng AS, Pegg TJ, Karamitsos TD, et al. Cardiovascular magnetic resonance perfusion imaging at 3-tesla for the detection of coronary artery disease: a comparison with 1.5-tesla. *J Am Coll Cardiol* 2007;49(25):2440-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Cheng CW, Yang NI, Lin KJ, et al. Role of coronary spasm for a positive noninvasive stress test result in angina pectoris patients without hemodynamically significant coronary artery disease. *Am J Med Sci* 2008;335(5):354-62. *Full-text exclusion reason(s): Data for women not reported as a subgroup; No outcomes of interest.*

Cheng V, Gutstein A, Wolak A, et al. Moving beyond binary grading of coronary arterial stenoses on coronary computed tomographic angiography: insights for the imager and referring clinician. *JACC Cardiovasc Imaging* 2008;1(4):460-71. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Cheng W, Zeng M, Arellano C, et al. Detection of myocardial perfusion abnormalities: Standard dual-source coronary computed tomography angiography versus rest/stress technetium-99m single-photo emission CT. *Br J Radiol* 2010;83(992):652-660. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Chikamori T, Doi YL, Yonezawa Y, et al. Noninvasive identification of significant narrowing of the left main coronary artery by dipyridamole thallium scintigraphy. *Am J Cardiol* 1991;68(5):472-7. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Chikamori T, Hirakawa K, Seo H, et al. Diagnostic significance of exercise-induced ST-segment depression in the lateral limb leads in patients with suspected coronary artery disease. *Am J Cardiol* 1995;76(7):513-6. *Full-text*

*exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Chiou KR, Huang WC, Lin SL, et al. Real-time dobutamine stress myocardial contrast echocardiography for detecting coronary artery disease: correlating abnormal wall motion and disturbed perfusion. *Can J Cardiol* 2004;20(12):1237-43. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Cho JS, Youn HJ, Cho EJ, et al. Dyssynchrony contributes to false-positive myocardial perfusion SPECT results in patients with stable angina. *Eur J Echocardiogr* 2011;12(6):461-6. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Choi EK, Chun EJ, Choi SI, et al. Assessment of subclinical coronary atherosclerosis in asymptomatic patients with type 2 diabetes mellitus with single photon emission computed tomography and coronary computed tomography angiography. *Am J Cardiol* 2009;104(7):890-6. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Choi JO, Cho SW, Song YB, et al. Longitudinal 2D strain at rest predicts the presence of left main and three vessel coronary artery disease in patients without regional wall motion abnormality. *Eur J Echocardiogr* 2009;10(5):695-701. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Choi SW, Cho KI, Lee HG, et al. Diagnostic value of ultrasound-based strain imaging in patients with suspected coronary artery disease. *Korean Circulation Journal* 2008;38(8):398-404. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Choi YS, Youn HJ, Park CS, et al. High echogenic thickening of proximal coronary artery predicts the far advanced coronary atherosclerosis. *Echocardiography* 2009;26(2):133-9. *Full-text exclusion reason(s): No data for NITs of interest (ECG, ECHO, SPECT, PET, CMR, CTA).*

Chotenimitkhun R, Hundley WG. Pharmacological stress cardiovascular magnetic resonance. *Postgrad Med* 2011;123(3):162-70. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Choudhary G, Shin V, Punjani S, et al. The role of calcium score and CT angiography in the medical management of patients with normal myocardial perfusion imaging. *J Nucl Cardiol* 2010;17(1):45-51. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Chow BJ, Abraham A, Wells GA, et al. Diagnostic accuracy and impact of computed tomographic coronary angiography on utilization of invasive coronary angiography. *Circ Cardiovasc Imaging* 2009;2(1):16-23. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Chow BJ, Al Shammeri OM, Beanlands RS, et al. Prognostic value of treadmill exercise and dobutamine stress positron emission tomography. *Can J Cardiol* 2009;25(7):e220-4. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Chow BJ, Ananthasubramaniam K, deKemp RA, et al. Comparison of treadmill exercise versus dipyridamole stress with myocardial perfusion imaging using rubidium-82 positron emission tomography. *J Am Coll Cardiol* 2005;45(8):1227-34. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Chow BJ, Dennie C, Hoffmann U, et al. Comparison of computed tomographic angiography versus rubidium-82 positron emission tomography for the detection of patients with anatomical coronary artery disease. *Can J Cardiol* 2007;23(10):801-7. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Chow BJ, Joseph P, Yam Y, et al. Usefulness of computed tomographic coronary angiography in patients with acute chest pain with and without high-risk features. *Am J Cardiol* 2010;106(4):463-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Chow BJ, Kass M, Gagne O, et al. Can differences in corrected coronary opacification measured with computed tomography predict resting coronary artery flow? *J Am Coll Cardiol* 2011;57(11):1280-8. *Full-text exclusion reason(s): Data for women not reported as a subgroup; No outcomes of interest.*

Chow BJW, Freeman MR, Bowen JM, et al. Ontario multidetector computed tomographic coronary angiography study: Field evaluation of diagnostic accuracy. *Arch Intern Med* 2011;171(11):1021-1029. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Choy JB, Leslie WD. Clinical correlates of Tc-99m sestamibi lung uptake. *J Nucl Cardiol* 2001;8(6):639-44. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Christensen HW, Haghfelt T, Vach W, et al. Observer reproducibility and validity of systems for clinical classification of angina pectoris: comparison with radionuclide imaging and coronary angiography. *Clin Physiol Funct Imaging* 2006;26(1):26-31. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Christian TF, Miller TD, Bailey KR, et al. Noninvasive identification of severe coronary artery disease using exercise tomographic thallium-201 imaging. *Am J Cardiol* 1992;70(1):14-20. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Christiansen JP, Edwards C, Sinclair T, et al. Detection of myocardial scar by contrast-enhanced cardiac magnetic resonance imaging in patients with troponin-positive chest pain and minimal angiographic coronary artery disease. *Am J Cardiol* 2006;97(6):768-71. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Chu ZG, Yang ZG, Dong ZH, et al. Characteristics of coronary artery disease in symptomatic type 2 diabetic patients: evaluation with CT angiography. *Cardiovasc Diabetol* 2010;9:74. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Chun EJ, Lee W, Choi YH, et al. Effects of nitroglycerin on the diagnostic accuracy of electrocardiogram-gated coronary computed tomography angiography. *J Comput Assist Tomogr* 2008;32(1):86-92. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Chung WY, Sir JJ, Cho YS, et al. Additive value of B-type natriuretic peptide on rest 201 Tl -dipyridamole stress 99m Tc -sestamibi gated myocardial SPECT in patients with normal left ventricular systolic function. *Cardiol Res Pract* 2010;1(1). *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Ciaroni S, Bloch A, Hoffmann JL, et al. Prognostic value of dobutamine echocardiography in patients with intermediate coronary lesions at angiography. *Echocardiography* 2002;19(7 Pt 1):549-53. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Cilli A, Batmaz F, Demir I, et al. The diagnostic yield of exercise stress testing as a screening tool for subclinical coronary artery disease in patients with moderate to severe obstructive sleep apnea. *J Clin Sleep Med* 2011;7(1):25-9. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Clouse ME, Sabir A, Yam CS, et al. Measuring noncalcified coronary atherosclerotic plaque using voxel analysis with MDCT angiography: a pilot clinical study. *AJR Am J Roentgenol* 2008;190(6):1553-60. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Coffey JP, Hill JC. Cardiac output and index in obese and non-obese patients using gated single photon emission computed tomography sestamibi perfusion imaging. *Journal of the Hong Kong College of Radiologists* 2005;8(4):226-232. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Cohen JL, Greene TO, Ottenweller J, et al. Dobutamine digital echocardiography for detecting coronary artery disease. *Am J Cardiol* 1991;67(16):1311-8. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Cohen JL, Ottenweller JE, George AK,, et al. Comparison of dobutamine and exercise echocardiography for detecting coronary artery disease. *Am J Cardiol* 1993;72(17):1226-31. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Cole JH, Chunn VM, Morrow JA,, et al. Cost implications of initial computed tomography angiography as opposed to catheterization in patients with mildly abnormal or equivocal myocardial perfusion scans. *J Cardiovasc Comput Tomogr* 2007;1(1):21-6. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup; No outcomes of interest.*

Coles DR, Smail MA, Negus IS,, et al. Comparison of radiation doses from multislice computed tomography coronary angiography and conventional diagnostic angiography. *J Am Coll Cardiol* 2006;47(9):1840-5. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Coles DR, Wilde P, Oberhoff M,, et al. Multislice computed tomography coronary angiography in patients admitted with a suspected acute coronary syndrome. *Int J Cardiovasc Imaging* 2007;23(5):603-14. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Coma-Canella I, Martinez-Caro D, Cosin-Sales J,, et al. Clandestine ischemia in patients with vasospastic angina. *Coron Artery Dis* 2000;11(5):383-90. *Full-text exclusion reason(s): Data for women not reported as a subgroup; No outcomes of interest.*

Coma-Canella I, Palazuelos J, Bravo N,, et al. Myocardial perfusion imaging with adenosine triphosphate predicts the rate of cardiovascular events. *J Nucl Cardiol* 2006;13(3):316-323. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Comess KA, Choi JH, Xie Z,, et al. Transthoracic coronary doppler vibrometry in the evaluation of normal volunteers and patients with coronary artery stenosis. *Ultrasound in Medicine and Biology* 2011;37(5):679-687. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Conti A, Sammiceli L, Gallini C,, et al. Assessment of patients with low-risk chest pain in the emergency department: Head-to-head comparison of exercise stress echocardiography and exercise myocardial SPECT. *Am Heart J* 2005;149(5):894-901. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Conti A, Vanni S, Sammiceli L,, et al. Yield of nuclear scan strategy in chest pain unit evaluation of special populations. *Nucl Med Commun* 2008;29(12):1106-12. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Conti A, Zanobetti M, Grifoni S,, et al. Implementation of myocardial perfusion imaging in the early triage of patients with suspected acute coronary syndromes. *Nucl Med Commun* 2003;24(10):1055-60. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Cortigiani L, Bigi R, Landi P,, et al. Prognostic implication of stress echocardiography in 6214 hypertensive and 5328 normotensive patients. *Euro Heart J* 2011;32(12):1509-1518. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Cortigiani L, Bigi R, Rigo F,, et al. Diagnostic value of exercise electrocardiography and dipyridamole stress echocardiography in hypertensive and normotensive chest pain patients with right bundle branch block. *J Hypertens* 2003;21(11):2189-94. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Cortigiani L, Coletta C, Bigi R,, et al. Clinical, exercise electrocardiographic, and pharmacologic stress echocardiographic findings for risk stratification of hypertensive patients with chest pain. *Am J Cardiol* 2003;91(8):941-5. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Cortigiani L, Rigo F, Gherardi S,, et al. Prognostic effect of coronary flow reserve in women versus men with chest pain syndrome and normal dipyridamole stress echocardiography. *Am J Cardiol* 2010;106(12):1703-8. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization.*

Cosson E, Guimfack M, Paries J,, et al. Are silent coronary stenoses predictable in diabetic patients and predictive of cardiovascular events? *Diabetes Metab* 2003;29(5):470-6. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Costa MA, Shoemaker S, Futamatsu H,, et al. Quantitative magnetic resonance perfusion imaging detects anatomic and physiologic coronary artery disease as measured by coronary angiography and fractional flow reserve. *J Am Coll Cardiol* 2007;50(6):514-22. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Cosyns B, Lancellotti P, Van Camp G,, et al. Head to head comparison of transesophageal and transthoracic contrast-enhanced echocardiography during dobutamine administration for the detection of coronary artery disease. *Int J Cardiol* 2008;129(1):105-10. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Coyne EP, Belvedere DA, Vande Streek PR,, et al. Thallium-201 scintigraphy after intravenous infusion of adenosine compared with exercise thallium testing in the diagnosis of coronary artery disease. *J Am Coll Cardiol* 1991;17(6):1289-94. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Cramer MJ, van der Wall EE, Jaarsma W,, et al. Detection of coronary artery disease: comparison between technetium 99m-labeled sestamibi single-photon emission computed tomography and two-dimensional echocardiography with dipyridamole low-level exercise-stress. *J Nucl Cardiol* 1996;3(5):389-94. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Crouse LJ, Harbrecht JJ, Vacek JL,, et al. Exercise echocardiography as a screening test for coronary artery disease and correlation with coronary arteriography. *Am J Cardiol* 1991;67(15):1213-8. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Cury RC, Cattani CA, Gabure LA,, et al. Diagnostic performance of stress perfusion and delayed-enhancement MR imaging in patients with coronary artery disease. *Radiology* 2006;240(1):39-45. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Cury RC, Ferencik M, Achenbach S,, et al. Accuracy of 16-slice multi-detector CT to quantify the degree of coronary artery stenosis: assessment of cross-sectional and longitudinal vessel reconstructions. *Eur J Radiol* 2006;57(3):345-50. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Cury RC, Magalhaes TA, Borges AC,, et al. Dipyridamole stress and rest myocardial perfusion by 64-detector row computed tomography in patients with suspected coronary artery disease. *Am J Cardiol* 2010;106(3):310-5. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Cury RC, Pomerantsev EV, Ferencik M,, et al. Comparison of the degree of coronary stenoses by multidetector computed tomography versus by quantitative coronary angiography. *Am J Cardiol* 2005;96(6):784-7. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup; No outcomes of interest.*

Cury RC, Techasith T, Feuchtner G,, et al. Cardiovascular disease and stroke in women: role of radiology. *AJR Am J Roentgenol* 2011;196(2):265-73. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Cwajg J, Xie F, O'Leary E,, et al. Detection of angiographically significant coronary artery disease with accelerated intermittent imaging after intravenous administration of ultrasound contrast material. *Am Heart J* 2000;139(4):675-83. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Dagianti A, Penco M, Agati L,, et al. Stress echocardiography: comparison of exercise, dipyridamole and dobutamine in detecting and predicting the extent of coronary artery disease. *J Am Coll Cardiol* 1995;26(1):18-25. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Dahan M, Viron BM, Poiseau E,, et al. Combined dipyridamole-exercise stress echocardiography for detection of myocardial ischemia in hemodialysis patients: an alternative to stress nuclear imaging. *Am J Kidney Dis* 2002;40(4):737-44. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Daimon M, Watanabe H, Yamagishi H,, et al. Physiologic assessment of coronary artery stenosis without stress tests: noninvasive analysis of phasic flow characteristics by transthoracic Doppler echocardiography. *J Am Soc Echocardiogr* 2005;18(9):949-55. *Full-text exclusion reason(s): No data for NITs of interest (ECG, ECHO, SPECT, PET, CMR, CTA).*

Daimon M, Watanabe H, Yamagishi H,, et al. Physiologic assessment of coronary artery stenosis by coronary flow reserve measurements with transthoracic Doppler echocardiography: comparison with exercise thallium-201 single piston emission computed tomography. *J Am Coll Cardiol* 2001;37(5):1310-5. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Dal Porto R, Faletta F, Picano E,, et al. Safety, feasibility, and diagnostic accuracy of accelerated high-dose dipyridamole stress echocardiography. *Am J Cardiol* 2001;87(5):520-4. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Danciu SC, Herrera CJ, Stecy PJ,, et al. Usefulness of multislice computed tomographic coronary angiography to identify patients with abnormal myocardial perfusion stress in whom diagnostic catheterization may be safely avoided. *Am J Cardiol* 2007;100(11):1605-8. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Danias PG, Ahlberg AW, Travin MI,, et al. Visual assessment of left ventricular perfusion and function with electrocardiography-gated SPECT has high intraobserver and interobserver reproducibility among experienced nuclear cardiologists and cardiology trainees. *J Nucl Cardiol* 2002;9(3):263-70. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; No outcomes of interest.*

Daou D, Delahaye N, Vilain D,, et al. Identification of extensive coronary artery disease: incremental value of exercise TI-201 SPECT to clinical and stress test variables. *J Nucl Cardiol* 2002;9(2):161-8. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Daoud EG, Pitt A, Armstrong WF. Electrocardiographic response during dobutamine stress echocardiography. *Am Heart J* 1995;129(4):672-7. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Dart J, Yuda S, Cain P,, et al. Use of myocardial backscatter as a quantitative tool for dobutamine echocardiography: feasibility, response to ischemia and accuracy compared with coronary angiography. *Int J Cardiovasc Imaging* 2002;18(5):325-36. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Dash H, Massie BM, Botvinick EH,, et al. The noninvasive identification of left main and three-vessel coronary artery disease by myocardial stress perfusion scintigraphy and treadmill exercise electrocardiography. *Circulation* 1979;60(2):276-84. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Dattilo G, Patane S, Zito C,, et al. Handgrip exercise associated with dobutamine stress echocardiography. *Int J Cardiol* 2010;143(3):298-301. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup; No outcomes of interest.*

Daugherty SL, Peterson PN, Magid DJ,, et al. The relationship between gender and clinical management after exercise stress testing. *Am Heart J* 2008;156(2):301-7. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization.*

Davar J, Roberts EB, Coghlan JG,, et al. Pacing-dobutamine stress echocardiography is a method of choice for non-invasive diagnosis of coronary artery disease in selected patients. *Eurecho 4, Lisbon (Portugal), 6-9 Dec 2000 (World Meeting Number 000 5377) 2000.* *Full-text exclusion reason(s): Conference abstract or trial registry posting.*

Davin L, Lancellotti P, Bruyere PJ,, et al. Diagnostic accuracy of computed tomography coronary angiography in routine practice. *Acta Cardiol* 2007;62(4):339-44. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Dawson D, Kaul S, Peters D,, et al. Prognostic value of dipyridamole stress myocardial contrast echocardiography: comparison with single photon emission computed tomography. *J Am Soc Echocardiogr* 2009;22(8):954-60. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for*

*symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Dawson D, Rinkevich D, Belcik T, et al. Measurement of myocardial blood flow velocity reserve with myocardial contrast echocardiography in patients with suspected coronary artery disease: comparison with quantitative gated Technetium 99m sestamibi single photon emission computed tomography. *J Am Soc Echocardiogr* 2003;16(11):1171-7. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup; No outcomes of interest.*

de Albuquerque Fonseca L, Picano E. Comparison of dipyridamole and exercise stress echocardiography for detection of coronary artery disease (a meta-analysis). *Am J Cardiol* 2001;87(10):1193-6; A4. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

De Almeida MC, Markman Filho B. Prognostic value of dipyridamole stress echocardiography in women. *Arq Bras Cardiol* 2011;96(1):31-37. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization.*

De Bello V, Bellina CR, Molea N, et al. Simultaneous dobutamine stress echocardiography and dobutamine scintigraphy (99mTc-MIBI-SPET) for assessment of coronary artery disease. *Int J Card Imaging* 1996;12(3):185-90. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

de Feyter PJ. Can multislice CT detect coronary artery disease accurately? *Nat Clin Pract Cardiovasc Med* 2005;2(11):560-1. *Full-text exclusion reason(s): Full-text unobtainable.*

de Graaf FR, Schuijf JD, Scholte AJ, et al. Usefulness of hypertriglyceridemic waist phenotype in type 2 diabetes mellitus to predict the presence of coronary artery disease as assessed by computed tomographic coronary angiography. *Am J Cardiol* 2010;106(12):1747-53. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

De Graaf FR, Schuijf JD, Van Velzen JE, et al. Diagnostic accuracy of 320-row multidetector computed tomography coronary angiography in the non-invasive evaluation of significant coronary artery disease. *Euro Heart J* 2010;31(15):1908-1915. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

De Graaf FR, Schuijf JD, Van Velzen JE, et al. Assessment of global left ventricular function and volumes with 320-row multidetector computed tomography: A comparison with 2D-echocardiography. *J Nucl Cardiol* 2010;17(2):225-231. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup; No outcomes of interest.*

de Leon G, Aguade-Bruix S, Aliaga V, et al. Submaximal exercise testing plus atropine in myocardial perfusion SPECT. *Rev Esp Cardiol* 2010;63(10):1155-61. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

De Lima JJ, Sabbaga E, Vieira ML, et al. Coronary angiography is the best predictor of events in renal transplant candidates compared with noninvasive testing. *Hypertension* 2003;42(3):263-8. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

De Lorenzo A, Hachamovitch R, Kang X, et al. Prognostic value of myocardial perfusion SPECT versus exercise electrocardiography in patients with ST-segment depression on resting electrocardiography. *J Nucl Cardiol* 2005;12(6):655-61. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup; No outcomes of interest.*

de Vries J, DeJongste MJ, Jessurun GA, et al. Myocardial perfusion quantification in patients suspected of cardiac syndrome X with positive and negative exercise testing: a [<sup>13</sup>N]ammonia positron emission tomography study. *Nucl Med Commun* 2006;27(10):791-4. *Full-text exclusion reason(s): Data for women not reported as a subgroup; No outcomes of interest.*

De Vries ST, Kleijn SA, Van THAWJ, et al. Impact of high altitude on echocardiographically determined cardiac morphology and function in patients with coronary artery disease and healthy controls. *European Journal of*



Echocardiography 2010;11(5):446-450. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup; No outcomes of interest.*

DeCara JM. Noninvasive cardiac testing in women. J Am Med Womens Assoc 2003;58(4):254-63. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Deetjen A, Mollmann S, Conradi G, et al. Use of automatic exposure control in multislice computed tomography of the coronaries: comparison of 16-slice and 64-slice scanner data with conventional coronary angiography. Heart 2007;93(9):1040-3. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Deetjen AG, Conradi G, Mollmann S, et al. Diagnostic value of the 16-detector row multislice spiral computed tomography for the detection of coronary artery stenosis in comparison to invasive coronary angiography. Clin Cardiol 2007;30(3):118-23. *Full-text exclusion reason(s): No outcomes of interest.*

deFilippi CR, Rosanio S, Tocchi M, et al. Randomized comparison of a strategy of predischarge coronary angiography versus exercise testing in low-risk patients in a chest pain unit: in-hospital and long-term outcomes. J Am Coll Cardiol 2001;37(8):2042-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Del Val Gomez M, Gallardo FG, Salazar ML, et al. [Prognostic value of normal myocardial radionuclide scan in patients with positive treadmill test]. Rev Esp Cardiol 2002;55(9):991-4. *Full-text exclusion reason(s): Non-English.*

Delcour KS, Khaja A, Chockalingam A, et al. Outcomes in patients with abnormal myocardial perfusion imaging and normal coronary angiogram. Angiology 2009;60(3):318-21. *Full-text exclusion reason(s): Population does not include women  $\geq$  age 18.*

Demer LL, Gould KL, Goldstein RA, et al. Assessment of coronary artery disease severity by positron emission tomography. Comparison with quantitative arteriography in 193 patients. Circulation 1989;79(4):825-35. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Demir H, Erbay G, Kir KM, et al. Clinical validation of technetium-99m MIBI-gated single-photon emission computed tomography (SPECT) for avoiding false positive results in patients with left bundle-branch block: comparison with stress-rest nongated SPECT. Clin Cardiol 2003;26(4):182-7. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup; No outcomes of interest.*

Dendukuri N, Chiu K, Brophy JM. Validity of electron beam computed tomography for coronary artery disease: asystematic review and meta-analysis. BMC Med 2007;5:35. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

DePuey EG, Guertler-Krawczynska E, D'Amato PH. Thallium-201 single photon emission computed tomography with intravenous dipyridamole to diagnose coronary artery disease. Coron Artery Dis 1990;75-82. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Derumeaux G, Redonnet M, Mouton-Schleifer D, et al. Dobutamine stress echocardiography in orthotopic heart transplant recipients. VACOMED Research Group. J Am Coll Cardiol 1995;25(7):1665-72. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Detrano R, Gianrossi R, Froelicher V. The diagnostic accuracy of the exercise electrocardiogram: a meta-analysis of 22 years of research. Prog Cardiovasc Dis 1989;32(3):173-206. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Detrano R, Hsiai T, Wang S, et al. Prognostic value of coronary calcification and angiographic stenoses in patients undergoing coronary angiography. J Am Coll Cardiol 1996;27(2):285-90. *Full-text exclusion reason(s): No data for NITs of interest (ECG, ECHO, SPECT, PET, CMR, CTA).*

Deutsch HJ, Schenkel C, Klaer R, et al. [Comparison of ergometer and dipyridamole echocardiography in patients with suspected coronary heart disease]. Z Kardiol 1994;83(6):446-53. *Full-text exclusion reason(s): Non-English.*

Dewey M, Dubel HP, Schink T,, et al. Head-to-head comparison of multislice computed tomography and exercise electrocardiography for diagnosis of coronary artery disease. *Eur Heart J* 2007;28(20):2485-90. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Dewey M, Hamm B. Cost effectiveness of coronary angiography and calcium scoring using CT and stress MRI for diagnosis of coronary artery disease. *Eur Radiol* 2007;17(5):1301-9. *Full-text exclusion reason(s): Not a clinical study report.*

Dewey M, Rutsch W, Schnapauff D,, et al. Coronary artery stenosis quantification using multislice computed tomography. *Invest Radiol* 2007;42(2):78-84. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Dewey M, Schnapauff D, Laule M,, et al. Multislice CT coronary angiography: evaluation of an automatic vessel detection tool. *Rofo* 2004;176(4):478-83. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup; No outcomes of interest.*

Dewey M, Teige F, Rutsch W,, et al. CT coronary angiography: influence of different cardiac reconstruction intervals on image quality and diagnostic accuracy. *Eur J Radiol* 2008;67(1):92-9. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Dewey M, Zimmermann E, Deissenrieder F,, et al. Noninvasive coronary angiography by 320-row computed tomography with lower radiation exposure and maintained diagnostic accuracy: comparison of results with cardiac catheterization in a head-to-head pilot investigation. *Circulation* 2009;120(10):867-75. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Di Bello V, Bellina CR, Gori E,, et al. Incremental diagnostic value of dobutamine stress echocardiography and dobutamine scintigraphy (technetium 99m-labeled sestamibi single-photon emission computed tomography) for assessment of presence and extent of coronary artery disease. *J Nucl Cardiol* 1996;3(3):212-20. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Di Bello V, Bellina CR, Molea N,, et al. Simultaneous dobutamine stress echocardiography and dobutamine scintigraphy (99mTc-MIBI-SPET) for assessment of coronary artery disease. *Int J Card Imaging* 1996;12(3):185-90. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Di Carli MF, Dorbala S. Cardiac PET-CT. *J Thorac Imaging* 2007;22(1):101-106. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Di Carli MF, Dorbala S, Curillova Z,, et al. Relationship between CT coronary angiography and stress perfusion imaging in patients with suspected ischemic heart disease assessed by integrated PET-CT imaging. *J Nucl Cardiol* 2007;14(6):799-809. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Di Cesare E, Battisti S, Riva A,, et al. Parallel imaging and dobutamine stress magnetic resonance imaging in patients with atypical chest pain or equivocal ECG not suitable for stress echocardiography. *Radiol Med* 2009;114(2):216-28. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Di Pasquale P, Cannizzaro S, Scalzo S,, et al. Sensitivity, specificity and predictive value of the echocardiography and troponin-T test combination in patients with non-ST elevation acute coronary syndromes. *Int J Cardiovasc Imaging* 2004;20(1):37-46. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup; No outcomes of interest.*

Di Pasquale P, Paterna S. Utility of the combination of echocardiography and troponin T test in patients with ST segment depression. *HeartDrug* 2002;2(6):279-285. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Diederichsen AC, Petersen H, Jensen LO, et al. Diagnostic value of cardiac 64-slice computed tomography: importance of coronary calcium. *Scand Cardiovasc J* 2009;43(5):337-44. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Diederichsen ACP, Petersen H, Jensen LO, et al. Diagnostic value of cardiac 64-slice computed tomography: Importance of coronary calcium. *Scand Cardiovasc J* 2009;43(5):337-344. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Dill T, Deetjen A, Ekinici O, et al. Radiation dose exposure in multislice computed tomography of the coronaries in comparison with conventional coronary angiography. *Int J Cardiol* 2008;124(3):307-11. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Dirksen MS, Jukema JW, Bax JJ, et al. Cardiac multidetector-row computed tomography in patients with unstable angina. *Am J Cardiol* 2005;95(4):457-61. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Djordjevic-Dikic AD, Ostojic MC, Beleslin BD, et al. High dose adenosine stress echocardiography for noninvasive detection of coronary artery disease. *J Am Coll Cardiol* 1996;28(7):1689-95. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Dodd JD, Rieber J, Pomerantsev E, et al. Quantification of nonculprit coronary lesions: comparison of cardiac 64-MDCT and invasive coronary angiography. *AJR Am J Roentgenol* 2008;191(2):432-8. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Dodla S, Xie F, Smith M, et al. Real-time perfusion echocardiography during treadmill exercise and dobutamine stress testing. *Heart* 2010;96(3):220-5. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Doesch C, Seeger A, Hoevelborn T, et al. Adenosine stress cardiac magnetic resonance imaging for the assessment of ischemic heart disease. *Clin Res Cardiol* 2008;97(12):905-12. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Dolan MS, Riad K, El-Shafei A, et al. Effect of intravenous contrast for left ventricular opacification and border definition on sensitivity and specificity of dobutamine stress echocardiography compared with coronary angiography in technically difficult patients. *Am Heart J* 2001;142(5):908-15. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Donati OF, Scheffel H, Stolzmann P, et al. Combined cardiac CT and MRI for the comprehensive workup of hemodynamically relevant coronary stenoses. *AJR Am J Roentgenol* 2010;194(4):920-6. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Dong S, Liang X, Zhang S, et al. Assessment of coronary artery disease with second harmonic myocardial perfusion contrast echocardiography. *Chin Med J (Engl)* 2002;115(6):837-41. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Dorgelo J, Willems TP, Geluk CA, et al. Multidetector computed tomography-guided treatment strategy in patients with non-ST elevation acute coronary syndromes: a pilot study. *Eur Radiol* 2005;15(4):708-13. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Dou KF, Yang MF, Yang YJ, et al. Myocardial 18F-FDG uptake after exercise-induced myocardial ischemia in patients with coronary artery disease. *J Nucl Med* 2008;49(12):1986-91. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Douglas PS, Garcia MJ, Haines DE, et al. ACCF/AHA/ASA/ASNC/HFSA/HRS/SCAI/SCCM/SCCT/SCMR 2011 Appropriate Use Criteria for Echocardiography. A Report of the American College of Cardiology Foundation Appropriate Use Criteria Task Force, American Society of Echocardiography, American Heart Association, American Society of Nuclear Cardiology, Heart Failure Society of America, Heart Rhythm Society, Society for

Cardiovascular Angiography and Interventions, Society of Critical Care Medicine, Society of Cardiovascular Computed Tomography, and Society for Cardiovascular Magnetic Resonance Endorsed by the American College of Chest Physicians. *J Am Coll Cardiol* 2011;57(9):1126-66. *Full-text exclusion reason(s): Not a clinical study report.*

Downs SH, Black N. The feasibility of creating a checklist for the assessment of the methodological quality both of randomised and non-randomised studies of health care interventions. *Journal of Epidemiology and Community Health* 1998;52(6):377-84. *Full-text exclusion reason(s): No data for NITs of interest (ECG, ECHO, SPECT, PET, CMR, CTA).*

Dragu R, Kerner A, Gruberg L., et al. Angiographically uncertain left main coronary artery narrowings: correlation with multidetector computed tomography and intravascular ultrasound. *Int J Cardiovasc Imaging* 2008;24(5):557-63. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Dragu R, Rispler S, Ghersin E., et al. Contrast enhanced multi-detector computed tomography coronary angiography versus conventional invasive quantitative coronary angiography in acute coronary syndrome patients-correlation and bias. *Acute Card Care* 2006;8(2):99-104. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Drosch T, Tsiflikas I, Brodoefel H., et al. Semi-automatic assessment of global left ventricular function and left ventricular parameters with dual-source computed tomography: comparison with invasive angiography. *Heart Vessels* 2010;25(1):57-62. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup; No outcomes of interest.*

Duarte PS, Mastrocolla LE, Sampaio CR., et al. Indication of myocardial perfusion scintigraphy for coronary artery disease detection based on clinical-epidemiological and treadmill test evidences. *Arq Bras Cardiol* 2006;87(4):415-22. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Duarte R, Bettencourt N, Costa JC., et al. Coronary computed tomography angiography in a single cardiac cycle with a mean radiation dose of approximately 1 mSv: initial experience. *Rev Port Cardiol* 2010;29(11):1667-76. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Dubart AE, Carvalho da Silva KG, Jr., Korosoglou G., et al. Real-time myocardial contrast echocardiography for the detection of stress-induced myocardial ischemia. Comparison with 99mTc-sestamibi single photon emission computed tomography. *Z Kardiol* 2004;93(11):890-6. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Dunn RF, Kelly DT, Bailey IK., et al. Serial exercise thallium myocardial perfusion scanning and exercise electrocardiography in the diagnosis of coronary artery disease. *Aust N Z J Med* 1979;9(5):547-53. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Duvall WL, Croft LB, Corriel JS., et al. SPECT myocardial perfusion imaging in morbidly obese patients: image quality, hemodynamic response to pharmacologic stress, and diagnostic and prognostic value. *J Nucl Cardiol* 2006;13(2):202-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Ebersole DG, Heironimus J, Toney MO., et al. Comparison of exercise and adenosine technetium-99m sestamibi myocardial scintigraphy for diagnosis of coronary artery disease in patients with left bundle branch block. *Am J Cardiol* 1993;71(5):450-3. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Eek C, Grenne B, Brunvand H., et al. Strain echocardiography predicts acute coronary occlusion in patients with non-ST-segment elevation acute coronary syndrome. *European Journal of Echocardiography* 2010;11(6):501-508. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Egger M, Juni P, Bartlett C., et al. How important are comprehensive literature searches and the assessment of trial quality in systematic reviews? Empirical study. *Health Technol Assess* 2003;7(1):1-76. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Ehara M, Surmely JF, Kawai M., et al. Diagnostic accuracy of 64-slice computed tomography for detecting angiographically significant coronary artery stenosis in an unselected consecutive patient population: comparison

with conventional invasive angiography. *Circ J* 2006;70(5):564-71. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Eisenberg MJ, Afilalo J, Lawler PR, et al. Cancer risk related to low-dose ionizing radiation from cardiac imaging in patients after acute myocardial infarction. *CMAJ* 2011;183(4):430-6. *Full-text exclusion reason(s): All women in the study are known to have CAD.*

Elamin MS, Boyle R, Kardash MM, et al. Accurate detection of coronary heart disease by new exercise test. *Br Heart J* 1982;48(4):311-20. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Elhendy A, Bax JJ, van Domburg RT, et al. Dobutamine stress thallium-201 single-photon emission tomography versus echocardiography for evaluation of the extent and location of coronary artery disease late after myocardial infarction. *Eur J Nucl Med* 1999;26(5):467-73. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Elhendy A, Geleijnse L, Salustri A, et al. T wave normalization during dobutamine stress testing in patients with non-Q wave myocardial infarction. A marker of myocardial ischaemia? *Eur Heart J* 1996;17(4):526-31. *Full-text exclusion reason(s): All women in the study are known to have CAD.*

Elhendy A, Geleijnse ML, Roelandt JR, et al. Comparison of dobutamine stress echocardiography and 99m-technetium sestamibi SPECT myocardial perfusion scintigraphy for predicting extent of coronary artery disease in patients with healed myocardial infarction. *Am J Cardiol* 1997;79(1):7-12. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Elhendy A, Geleijnse ML, van Domburg RT, et al. Comparison of dobutamine stress echocardiography and technetium-99m sestamibi single-photon emission tomography for the diagnosis of coronary artery disease in hypertensive patients with and without left ventricular hypertrophy. *Eur J Nucl Med* 1998;25(1):69-78. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Elhendy A, O'Leary EL, Xie F, et al. Comparative accuracy of real-time myocardial contrast perfusion imaging and wall motion analysis during dobutamine stress echocardiography for the diagnosis of coronary artery disease. *J Am Coll Cardiol* 2004;44(11):2185-91. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Elhendy A, Roelandt J, Geleijnse ML, et al. Non-invasive diagnosis of coronary artery disease in hypertensive patients. Comparison of dobutamine stress echocardiography and technetium sestamibi SPECT imaging. XIXth Congress of the European Society of Cardiology, Stockholm (Sweden), 24-28 August 1997 (World Meeting Number 973 0006); 1997.

Elhendy A, van Domburg RT, Bax JJ, et al. Optimal criteria for the diagnosis of coronary artery disease by dobutamine stress echocardiography. *Am J Cardiol* 1998;82(11):1339-44. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Elhendy A, van Domburg RT, Bax JJ, et al. Accuracy of dobutamine technetium 99m sestamibi SPECT imaging for the diagnosis of single-vessel coronary artery disease: comparison with echocardiography. *Am Heart J* 2000;139(2 Pt 1):224-30. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Elhendy A, van Domburg RT, Bax JJ, et al. Significance of resting wall motion abnormalities in 2-dimensional echocardiography in patients without previous myocardial infarction referred for pharmacologic stress testing. *J Am Soc Echocardiogr* 2000;13(1):1-8. *Full-text exclusion reason(s): Data for women not reported as a subgroup; No outcomes of interest.*

Elhendy A, van Domburg RT, Poldermans D, et al. Safety and feasibility of dobutamine-atropine stress echocardiography for the diagnosis of coronary artery disease in diabetic patients unable to perform an exercise stress test. *Diabetes Care* 1998;21(11):1797-802. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Elhendy A, van Domburg RT, Roelandt JR,, et al. Safety and feasibility of dobutamine-atropine stress testing in hypertensive patients. *Hypertension* 1997;29(6):1232-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Elhendy A, van Domburg RT, Sozzi FB,, et al. Impact of hypertension on the accuracy of exercise stress myocardial perfusion imaging for the diagnosis of coronary artery disease. *Heart* 2001;85(6):655-61. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Elhendy A, yan Domburg RT, Roelandt JR,, et al. Accuracy of dobutamine stress echocardiography for the diagnosis of coronary artery stenosis in patients with myocardial infarction: the impact of extent and severity of left ventricular dysfunction. *Heart* 1996;76(2):123-8. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Ellestad MH, Crump R, Surber M. The significance of lead strength on ST changes during treadmill stress tests. *J Electrocardiol* 1992;25 Suppl:31-4. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

El-Mahalawy N, Abdel-Salam Z, Samir A,, et al. Left ventricular transient ischemic dilation during dobutamine stress echocardiography predicts multi-vessel coronary artery disease. *J Cardiol* 2009;54(2):255-61. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Emre A, Ersek B, Gursurer M,, et al. Angiographic and scintigraphic (perfusion and electrocardiogram-gated SPECT) correlates of clinical presentation in unstable angina. *Clin Cardiol* 2000;23(7):495-500. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup; No outcomes of interest.*

Epstein M, Gin K, Sterns L,, et al. Dobutamine stress echocardiography: initial experience of a Canadian centre. *Can J Cardiol* 1992;8(3):273-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Erbel R, Budde T, Kerkhoff G,, et al. Understanding the pathophysiology of the arterial wall: Which method should we choose? Electron beam computed tomography. *Euro Heart J* 2002;4(6). *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Erdogan N, Akar N, Vural M,, et al. Diagnostic value of 16-slice multidetector computed tomography in symptomatic patients with suspected significant obstructive coronary artery disease. *Heart Vessels* 2006;21(5):278-84. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Eren S, Bayram E, Fil F,, et al. An investigation of the association between coronary artery dominance and coronary artery variations with coronary arterial disease by multidetector computed tomographic coronary angiography. *J Comput Assist Tomogr* 2008;32(6):929-33. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup; No outcomes of interest.*

Erhard I, Rieber J, Jung P,, et al. The validation of fractional flow reserve in patients with coronary multivessel disease: a comparison with SPECT and contrast-enhanced dobutamine stress echocardiography. *Z Kardiol* 2005;94(5):321-7. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup; No outcomes of interest.*

Eriksson SV. Vectorcardiography: A tool for non-invasive detection of reperfusion and reocclusion? *Thromb Haemost* 1999;82(SUPPL. 1):64-67. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Eroglu E, Bayrak F, Gemici G,, et al. Prevalence of coronary artery disease in low to moderate-risk asymptomatic women: a multislice computed tomography study. *Turk Kardiyol Dern Ars* 2008;36(7):439-45. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup.*

Eroglu E, D'Hooge J, Herbots L,, et al. Comparison of real-time tri-plane and conventional 2D dobutamine stress echocardiography for the assessment of coronary artery disease. *Eur Heart J* 2006;27(14):1719-24. *Full-text exclusion reason(s): Data for women not reported as a subgroup; No outcomes of interest.*

Eskola MJ, Nikus KC, Holmvang L,, et al. Value of the 12-lead electrocardiogram to define the level of obstruction in acute anterior wall myocardial infarction: correlation to coronary angiography and clinical outcome in the

DANAMI-2 trial. *Int J Cardiol* 2009(3):378-83. *Full-text exclusion reason(s): No data for NITs of interest (ECG, ECHO, SPECT, PET, CMR, CTA).*

Esteves FP, Khan A, Correia LCL, et al. Absent coronary artery calcium excludes inducible myocardial ischemia on computed tomography/positron emission tomography. *Int J Cardiol* 2011;147(3):424-427. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Esteves FP, Sanyal R, Nye JA, et al. Adenosine stress rubidium-82 PET/computed tomography in patients with known and suspected coronary artery disease. *Nucl Med Commun* 2008;29(8):674-8. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Esteves FP, Sanyal R, Santana CA, et al. Potential impact of noncontrast computed tomography as gatekeeper for myocardial perfusion positron emission tomography in patients admitted to the chest pain unit. *Am J Cardiol* 2008;101(2):149-52. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Fallahi B, Beiki D, Fard-Esfahani A, et al. The additive value of transient left ventricular dilation using two-day dipyridamole 99mTc-MIBI SPET for screening coronary artery disease in patients with otherwise normal myocardial perfusion: a comparison between diabetic and non-diabetic cases. *Hell J Nucl Med* 2010;13(3):246-52. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Fallahi B, Beiki D, Gholamrezanezhad A, et al. Single Tc99m Sestamibi injection, double acquisition gated SPECT after stress and during low-dose dobutamine infusion: a new suggested protocol for evaluation of myocardial perfusion. *Int J Cardiovasc Imaging* 2008;24(8):825-35. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Fang XM, Chen HW, Hu XY, et al. Dual-source CT coronary angiography without heart rate or rhythm control in comparison with conventional coronary angiography. *Int J Cardiovasc Imaging* 2010;26(3):323-31. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Farzaneh-Far A, Kwong RY. Detecting acute coronary syndromes by magnetic resonance imaging. *Heart and Metabolism* 2011(50):15-19. *Full-text exclusion reason(s): Not a clinical study report; Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry; Data for women not reported as a subgroup.*

Fazel P, Peterman MA, Schussler JM. Three-year outcomes and cost analysis in patients receiving 64-slice computed tomographic coronary angiography for chest pain. *Am J Cardiol* 2009;104(4):498-500. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Feldman C, Vitola D, Schiavo N. Detection of coronary artery disease based on the calcification index obtained by helical computed tomography. *Arq Bras Cardiol* 2000;75(6):471-80. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Feldman JA, Bernard S, Mitchell P, et al. Effects of cardiology review of the electrocardiogram in patients with suspected acute coronary syndromes. *Am J Emerg Med* 2011;29(3):309-15 e2. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Felsher J, Meissner MD, Hakki AH, et al. Exercise thallium imaging in patients with diabetes mellitus. Prognostic implications. *Arch Intern Med* 1987;147(2):313-7. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Fenchel M, Franow A, Martirosian P, et al. 1 M Gd-chelate (gadobutrol) for multislice first-pass magnetic resonance myocardial perfusion imaging. *Br J Radiol* 2007;80(959):884-92. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Feola M, Biggi A, Ribichini F, et al. The diagnosis of coronary artery disease in hypertensive patients with chest pain and complete left bundle branch block: utility of adenosine Tc-99m tetrofosmin SPECT. *Clin Nucl Med* 2002;27(7):510-5. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Feola M, Biggi A, Vado A,, et al. The usefulness of adenosine 99mTc tetrofosmin SPECT for the diagnosis of left anterior descending coronary artery disease in patients with chest pain and left bundle branch block. *Nucl Med Commun* 2004;25(3):265-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Ferencik M, Nomura CH, Maurovich-Horvat P,, et al. Quantitative parameters of image quality in 64-slice computed tomography angiography of the coronary arteries. *Eur J Radiol* 2006;57(3):373-9. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup; No outcomes of interest.*

Ferencik M, Ropers D, Abbara S,, et al. Diagnostic accuracy of image postprocessing methods for the detection of coronary artery stenoses by using multidetector CT. *Radiology* 2007;243(3):696-702. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup; No outcomes of interest.*

Fernandes LP, Tardif JC, Arsenault A,, et al. Detection of myocardial perfusion abnormalities after a recent acute coronary syndrome by quantitative Levovist myocardial contrast echocardiography: comparison with 99m Tc-Myoview SPECT imaging. *Can J Cardiol* 2003;19(3):251-6. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Fernandez-Friera L, Garcia-Alvarez A, Bagheriannejad-Esfahani F,, et al. Diagnostic value of coronary artery calcium scoring in low-intermediate risk patients evaluated in the emergency department for acute coronary syndrome. *Am J Cardiol* 2011;107(1):17-23. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Ferrara N, Leosco D, Abete P,, et al. Dipyridamole echocardiography as a useful and safe test in the assessment of coronary artery disease in the elderly. *J Am Geriatr Soc* 1991;39(10):993-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Fesmire FM, Hughes AD, Stout PK,, et al. Selective dual nuclear scanning in low-risk patients with chest pain to reliably identify and exclude acute coronary syndromes. *Ann Emerg Med* 2001;38(3):207-15. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Feuchtner G, Postel T, Weidinger F,, et al. Is there a relation between non-calcifying coronary plaques and acute coronary syndromes? A retrospective study using multislice computed tomography. *Cardiology* 2008;110(4):241-8. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Feuchtner GM, Cury RC, Jodocy D,, et al. Differences in coronary plaque composition by noninvasive computed tomography angiography in individuals with and without obstructive coronary artery disease. *Atherosclerosis* 2011;215(1):90-5. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Fine JJ, Hopkins CB, Hall PA,, et al. Noninvasive coronary angiography: agreement of multi-slice spiral computed tomography and selective catheter angiography. *Int J Cardiovasc Imaging* 2004;20(6):549-52. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Fine JJ, Hopkins CB, Ruff N,, et al. Comparison of accuracy of 64-slice cardiovascular computed tomography with coronary angiography in patients with suspected coronary artery disease. *Am J Cardiol* 2006;97(2):173-4. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Finkelhor RS, Newhouse KE, Vrobel TR,, et al. The ST segment/heart rate slope as a predictor of coronary artery disease: comparison with quantitative thallium imaging and conventional ST segment criteria. *Am Heart J* 1986;112(2):296-304. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*



Fleischmann KE, Hunink MG, Kuntz KM,, et al. Exercise echocardiography or exercise SPECT imaging? A meta-analysis of diagnostic test performance. *JAMA* 1998;280(10):913-20. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Fleischmann S, Koepfli P, Namdar M,, et al. Gated (99m)Tc-tetrofosmin SPECT for discriminating infarct from artifact in fixed myocardial perfusion defects. *J Nucl Med* 2004;45(5):754-9. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Fleming RM. High-dose dipyridamole and gated sestamibi SPECT imaging provide diagnostic resting and stress ejection fractions useful for predicting extent of coronary artery disease. *Angiology* 2002;53(4):415-21. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Fleming RM, Kirkeeide RL, Taegtmeier H,, et al. Comparison of technetium-99m tetrofosmin tomography with automated quantitative coronary arteriography and thallium-201 tomographic imaging. *J Am Coll Cardiol* 1991;17(6):1297-302. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Flohr TG, McCollough CH, Bruder H,, et al. First performance evaluation of a dual-source CT (DSCT) system. *Eur Radiol* 2006;16(2):256-68. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup; No outcomes of interest.*

Flohr TG, Schoepf UJ, Kuettner A,, et al. Advances in cardiac imaging with 16-section CT systems. *Acad Radiol* 2003;10(4):386-401. *Full-text exclusion reason(s): Not a clinical study report.*

Florenciano-Sanchez R, de la Morena-Valenzuela G, Villegas-Garcia M,, et al. Noninvasive assessment of coronary flow velocity reserve in left anterior descending artery adds diagnostic value to both clinical variables and dobutamine echocardiography: a study based on clinical practice. *Eur J Echocardiogr* 2005;6(4):251-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Forberg JL, Hilmersson CE, Carlsson M,, et al. Negative predictive value and potential cost savings of acute nuclear myocardial perfusion imaging in low risk patients with suspected acute coronary syndrome: a prospective single blinded study. *BMC Emerg Med* 2009;9:12. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Forster S, Rieber J, Ubleis C,, et al. Tc-99m sestamibi single photon emission computed tomography for guiding percutaneous coronary intervention in patients with multivessel disease: a comparison with quantitative coronary angiography and fractional flow reserve. *Int J Cardiovasc Imaging* 2010;26(2):203-13. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Forster T, McNeill AJ, Salustri A,, et al. Simultaneous dobutamine stress echocardiography and technetium-99m isonitrite single-photon emission computed tomography in patients with suspected coronary artery disease. *J Am Coll Cardiol* 1993;21(7):1591-6. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Fragasso G, Lu C, Dabrowski P,, et al. Comparison of stress/rest myocardial perfusion tomography, dipyridamole and dobutamine stress echocardiography for the detection of coronary disease in hypertensive patients with chest pain and positive exercise test. *J Am Coll Cardiol* 1999;34(2):441-7. *Full-text exclusion reason(s): All women in the study are known to have CAD.*

Freedman N, Schechter D, Klein M,, et al. SPECT attenuation artifacts in normal and overweight persons: insights from a retrospective comparison of Rb-82 positron emission tomography and TI-201 SPECT myocardial perfusion imaging. *Clin Nucl Med* 2000;25(12):1019-23. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup.*

Freeman JA, Corbin M, Dunn M,, et al. Correlation of single lead telemetric treadmill exercise testing with coronary angiography. *J Electrocardiol* 1973;6(3):231-4. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Frohwein S, Klein JL, Lane A,, et al. Transesophageal dobutamine stress echocardiography in the evaluation of coronary artery disease. *J Am Coll Cardiol* 1995;25(4):823-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

From AM, Kane G, Bruce C,, et al. Characteristics and outcomes of patients with abnormal stress echocardiograms and angiographically mild coronary artery disease (<50% stenoses) or normal coronary arteries. *J Am Soc Echocardiogr* 2010;23(2):207-14. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Fujitaka K, Nakamura S, Sugiura T,, et al. Combined analysis of multislice computed tomography coronary angiography and stress-rest myocardial perfusion imaging in detecting patients with significant proximal coronary artery stenosis. *Nucl Med Commun* 2009;30(10):789-96. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Fukai T, Koyanagi S, Tashiro H,, et al. Adenosine triphosphate stress echocardiography in the detection of myocardial ischemia. *Am J Card Imaging* 1995;9(4):237-44. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Fukuoka S, Maeno M, Nakagawa S,, et al. Feasibility of myocardial dual-isotope perfusion imaging combined with gated single photon emission tomography for assessing coronary artery disease. *Nucl Med Commun* 2002;23(1):19-29. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Futamatsu H, Klassen C, Pilla M,, et al. Diagnostic accuracy of quantitative cardiac MRI evaluation compared to stress single-photon-emission computed tomography. *Int J Cardiovasc Imaging* 2008;24(3):293-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Futamatsu H, Wilke N, Klassen C,, et al. Evaluation of cardiac magnetic resonance imaging parameters to detect anatomically and hemodynamically significant coronary artery disease. *Am Heart J* 2007;154(2):298-305. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Gaddi O, Tortorella G, Picano E,, et al. Diagnostic and prognostic value of vasodilator stress echocardiography in asymptomatic Type 2 diabetic patients with positive exercise thallium scintigraphy: a pilot study. *Diabet Med* 1999;16(9):762-6. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Gaemperli O, Schepis T, Koepfli P,, et al. Accuracy of 64-slice CT angiography for the detection of functionally relevant coronary stenoses as assessed with myocardial perfusion SPECT. *Eur J Nucl Med Mol Imaging* 2007;34(8):1162-71. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Gaemperli O, Schepis T, Valenta I,, et al. Functionally relevant coronary artery disease: comparison of 64-section CT angiography with myocardial perfusion SPECT. *Radiology* 2008;248(2):414-23. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Gaemperli O, Valenta I, Schepis T,, et al. Coronary 64-slice CT angiography predicts outcome in patients with known or suspected coronary artery disease. *Eur Radiol* 2008;18(6):1162-73. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Gaibazzi N, Reverberi C, Badano L. Usefulness of contrast stress-echocardiography or exercise-electrocardiography to predict long-term acute coronary syndromes in patients presenting with chest pain without electrocardiographic abnormalities or 12-hour troponin elevation. *Am J Cardiol* 2011;107(2):161-7. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Gaibazzi N, Reverberi C, Squeri A,, et al. Contrast stress echocardiography for the diagnosis of coronary artery disease in patients with chest pain but without acute coronary syndrome: incremental value of myocardial perfusion. *J Am Soc Echocardiogr* 2009;22(4):404-10. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Gaibazzi N, Rigo F, Reverberi C. Detection of coronary artery disease by combined assessment of wall motion, myocardial perfusion and coronary flow reserve: a multiparametric contrast stress-echocardiography study. *J Am Soc Echocardiogr* 2010;23(12):1242-50. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Gaibazzi N, Rigo F, Squeri A,, et al. Incremental value of contrast myocardial perfusion to detect intermediate versus severe coronary artery stenosis during stress-echocardiography. *Cardiovasc Ultrasound* 2010;8:16. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Galanti G, Sciagra R, Comeglio M,, et al. Diagnostic accuracy of peak exercise echocardiography in coronary artery disease: comparison with thallium-201 myocardial scintigraphy. *Am Heart J* 1991;122(6):1609-16. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Galassi AR, Azzarelli S, Lupo L,, et al. Accuracy of exercise testing in the assessment of the severity of myocardial ischemia as determined by means of technetium-99m tetrofosmin SPECT scintigraphy. *J Nucl Cardiol* 2000;7(6):575-83. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Gallagher MJ, Ross MA, Raff GL,, et al. The diagnostic accuracy of 64-slice computed tomography coronary angiography compared with stress nuclear imaging in emergency department low-risk chest pain patients. *Ann Emerg Med* 2007;49(2):125-36. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Garber AM, Solomon NA. Cost-effectiveness of alternative test strategies for the diagnosis of coronary artery disease. *Ann Intern Med* 1999;130(9):719-28. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Garcia MJ, Lessick J, Hoffmann MH. Accuracy of 16-row multidetector computed tomography for the assessment of coronary artery stenosis. *JAMA* 2006;296(4):403-11. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Gaspar T, Dvir D, Peled N. The role of 16-slice computed tomography angiography in the diagnosis of coronary artery disease: large sample analysis. *Isr Med Assoc J* 2005;7(7):424-7. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Gaudio C, Mirabelli F, Alessandra L,, et al. Noninvasive assessment of coronary artery stenoses by multidetector-row spiral computed tomography: comparison with conventional angiography. *Eur Rev Med Pharmacol Sci* 2005;9(1):13-21. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Gaudio C, Tanzilli G, Vittore A,, et al. Detection of coronary artery stenoses using breath-hold magnetic resonance coronary angiography. Comparison with conventional x-ray angiography. *Eur Rev Med Pharmacol Sci* 2004;8(3):121-8. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Gebker R, Jahnke C, Manka R,, et al. Additional value of myocardial perfusion imaging during dobutamine stress magnetic resonance for the assessment of coronary artery disease. *Circ Cardiovasc Imaging* 2008;1(2):122-30. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Gebker R, Jahnke C, Paetsch I,, et al. Diagnostic performance of myocardial perfusion MR at 3 T in patients with coronary artery disease. *Radiology* 2008;247(1):57-63. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Geleijns J, Joemai RM, Dewey M,, et al. Radiation exposure to patients in a multicenter coronary angiography trial (CORE 64). *AJR Am J Roentgenol* 2011;196(5):1126-32. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Geleijnse ML, Elhendy A. Can stress echocardiography compete with perfusion scintigraphy in the detection of coronary artery disease and cardiac risk assessment? *Eur J Echocardiogr* 2000;1(1):12-21. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Geleijnse ML, Krenning BJ, Soliman OI,, et al. Dobutamine stress echocardiography for the detection of coronary artery disease in women. *Am J Cardiol* 2007;99(5):714-7. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Geleijnse ML, Marwick TH, Boersma E,, et al. Optimal pharmacological stress testing for the diagnosis of coronary artery disease: a probabilistic approach. *Eur Heart J* 1995;16 Suppl M:3-10. *Full-text exclusion reason(s): Not*

*original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry; Data for women not reported as a subgroup.*

Geleijnse ML, Vigna C, Kasprzak JD, et al. Usefulness and limitations of dobutamine-atropine stress echocardiography for the diagnosis of coronary artery disease in patients with left bundle branch block. A multicentre study. *Eur Heart J* 2000;21(20):1666-73. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Geluk CA, Dijkers R, Perik PJ, et al. Measurement of coronary calcium scores by electron beam computed tomography or exercise testing as initial diagnostic tool in low-risk patients with suspected coronary artery disease. *Eur Radiol* 2008;18(2):244-52. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Genovesi D, Giorgetti A, Gimelli A, et al. Impact of attenuation correction and gated acquisition in SPECT myocardial perfusion imaging: results of the multicentre SPAG (SPECT Attenuation Correction vs Gated) study. *Eur J Nucl Med Mol Imaging* 2011. *Full-text exclusion reason(s): No outcomes of interest.*

George RT, Arbab-Zadeh A, Miller JM, et al. Adenosine stress 64- and 256-row detector computed tomography angiography and perfusion imaging: a pilot study evaluating the transmural extent of perfusion abnormalities to predict atherosclerosis causing myocardial ischemia. *Circ Cardiovasc Imaging* 2009;2(3):174-82. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Gerber BL, Coche E, Pasquet A, et al. Coronary artery stenosis: direct comparison of four-section multi-detector row CT and 3D navigator MR imaging for detection—initial results. *Radiology* 2005;234(1):98-108. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Gerber TC, Sheedy PF, Bell MR, et al. Evaluation of the coronary venous system using electron beam computed tomography. *Int J Cardiovasc Imaging* 2001;17(1):65-75. *Full-text exclusion reason(s): No data for NITs of interest (ECG, ECHO, SPECT, PET, CMR, CTA).*

Ghadri JR, Pazhenkottil AP, Nkoulou RN, et al. Very high coronary calcium score unmasks obstructive coronary artery disease in patients with normal SPECT MPI. *Heart* 2011;97(12):998-1003. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Ghaffari S, Toufan M. The value of endothelium dependent vasodilatation in diagnosing coronary artery disease and its comparison with the results of routine diagnostic tests. *Saudi Med J* 2007;28(9):1344-9. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Ghersin E, Litmanovich D, Dragu R, et al. 16-MDCT coronary angiography versus invasive coronary angiography in acute chest pain syndrome: a blinded prospective study. *AJR Am J Roentgenol* 2006;186(1):177-84. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Ghostine S, Caussin C, Daoud B, et al. Non-invasive detection of coronary artery disease in patients with left bundle branch block using 64-slice computed tomography. *J Am Coll Cardiol* 2006;48(10):1929-34. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Ghostine S, Caussin C, Daoud B, et al. Noninvasive Detection of Coronary Artery Disease in Patients with Left Bundle Branch Block Using 64-Slice Computed Tomography. 55th Annual Scientific Session of the American College of Cardiology, Georgia World Congress Center, Atlanta, Georgia (USA), 11-14 Mar 2006 2006. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Giardina A, De Castro S, Fedele F, et al. Noninvasive testing for coronary artery disease in women. *Minerva Cardioangiol* 2006;54(3):323-30. *Full-text exclusion reason(s): Not a clinical study report.*

Gibbons RJ, Balady GJ, Beasley JW, et al. ACC/AHA Guidelines for Exercise Testing. A report of the American College of Cardiology/American Heart Association Task Force on Practice Guidelines (Committee on Exercise Testing). *J Am Coll Cardiol* 1997;30(1):260-311. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Gibson CM, Kirtane AJ, Murphy SA,, et al. Early initiation of eptifibatide in the emergency department before primary percutaneous coronary intervention for ST-segment elevation myocardial infarction: results of the Time to Integrilin Therapy in Acute Myocardial Infarction (TITAN)-TIMI 34 trial. *Am Heart J* 2006;152(4):668-75. *Full-text exclusion reason(s): No data for NITs of interest (ECG, ECHO, SPECT, PET, CMR, CTA).*

Gibson PB, Demus D, Noto R,, et al. Low event rate for stress-only perfusion imaging in patients evaluated for chest pain. *J Am Coll Cardiol* 2002;39(6):999-1004. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Gigli G, Cortigiani L, Vallebona A,, et al. Vasodilator stress echocardiography for risk stratification of medically stabilized unstable angina. *Eur J Echocardiogr* 2002;3(1):59-66. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Gill JB, Ruddy TD, Newell JB,, et al. Prognostic importance of thallium uptake by the lungs during exercise in coronary artery disease. *N Engl J Med* 1987;317(24):1486-9. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Ginzton LE, Conant R, Brizendine M,, et al. Exercise subcostal two-dimensional echocardiography: a new method of segmental wall motion analysis. *Am J Cardiol* 1984;53(6):805-11. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Giorgetti A, Marzullo P, Sambuceti G,, et al. Baseline/post-nitrate Tc-99m tetrofosmin mismatch for the assessment of myocardial viability in patients with severe left ventricular dysfunction: comparison with baseline Tc-99m tetrofosmin scintigraphy/FDG PET imaging. *J Nucl Cardiol* 2004;11(2):142-51. *Full-text exclusion reason(s): All women in the study are known to have CAD.*

Giorgetti A, Rossi M, Stanislao M,, et al. Feasibility and diagnostic accuracy of a gated SPECT early-imaging protocol: a multicenter study of the Myoview Imaging Optimization Group. *J Nucl Med* 2007;48(10):1670-5. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Girotra S, Keelan M, Weinstein AR,, et al. Relation of heart rate response to exercise with prognosis and atherosclerotic progression after coronary artery bypass grafting. *Am J Cardiol* 2009;103(10):1386-90. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Go RT, Marwick TH, MacIntyre WJ,, et al. A prospective comparison of rubidium-82 PET and thallium-201 SPECT myocardial perfusion imaging utilizing a single dipyridamole stress in the diagnosis of coronary artery disease. *J Nucl Med* 1990;31(12):1899-905. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Goldschlager N, Selzer A, Cohn K. Treadmill stress tests as indicators of presence and severity of coronary artery disease. *Ann Intern Med* 1976;85(3):277-86. *Full-text exclusion reason(s): Population does not include women  $\geq$  age 18; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Goldstein JA, Gallagher MJ, O'Neill WW,, et al. A randomized controlled trial of multi-slice coronary computed tomography for evaluation of acute chest pain. *J Am Coll Cardiol* 2007;49(8):863-71. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Gonzalez P, Massardo T, Jofre MJ,, et al. (201)Tl myocardial SPECT detects significant coronary artery disease between 50% and 75% angiogram stenosis. *Revista Espanola de Medicina Nuclear* 2005;24(5):305-311. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Gonzalez P, Massardo T, Jofre MJ,, et al. 201Tl myocardial SPECT detects significant coronary artery disease between 50% and 75% angiogram stenosis. *Rev Esp Med Nucl* 2005;24(5):305-11. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Gopal A, Nasir K, Ahmadi N, et al. Cardiac computed tomographic angiography in an outpatient setting: an analysis of clinical outcomes over a 40-month period. *J Cardiovasc Comput Tomogr* 2009;3(2):90-5. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Gottdiener JS. Overview of stress echocardiography: uses, advantages, and limitations. *Prog Cardiovasc Dis* 2001;43(4):315-34. *Full-text exclusion reason(s): Not a clinical study report.*

Gould KL. Positron emission tomography and interventional cardiology. *Am J Cardiol* 1990;66(14):51F-58F. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Gould KL, Goldstein RA, Mullani NA, et al. Noninvasive assessment of coronary stenoses by myocardial perfusion imaging during pharmacologic coronary vasodilation. VIII. Clinical feasibility of positron cardiac imaging without a cyclotron using generator-produced rubidium-82. *J Am Coll Cardiol* 1986;7(4):775-89. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Gouya H, Varenne O, Trinquart L, et al. Coronary artery stenosis in high-risk patients: 64-section CT and coronary angiography—prospective study and analysis of discordance. *Radiology* 2009;252(2):377-85. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Greenland P, Abrams J, Aurigemma GP, et al. Prevention Conference V: Beyond secondary prevention: identifying the high-risk patient for primary prevention: noninvasive tests of atherosclerotic burden: Writing Group III. *Circulation* 2000;101(1). *Full-text exclusion reason(s): Not a clinical study report.*

Greenland P, Alpert JS, Beller GA, et al. 2010 ACCF/AHA guideline for assessment of cardiovascular risk in asymptomatic adults: a report of the American College of Cardiology Foundation/American Heart Association Task Force on Practice Guidelines. *J Am Coll Cardiol* 2010;56(25):e50-103. *Full-text exclusion reason(s): Not a clinical study report.*

Greenland P, Smith SC, Jr., Grundy SM. Improving coronary heart disease risk assessment in asymptomatic people: role of traditional risk factors and noninvasive cardiovascular tests. *Circulation* 2001;104(15):1863-7. *Full-text exclusion reason(s): Not a clinical study report.*

Gregoire J, Theroux P. Detection and assessment of unstable angina using myocardial perfusion imaging: Comparison between technetium-99m sestamibi SPECT and 12-lead electrocardiogram. *Am J Cardiol* 1990;66(13):42E-46E. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Groothuis JG, Beek AM, Brinckman SL, et al. Low to intermediate probability of coronary artery disease: comparison of coronary CT angiography with first-pass MR myocardial perfusion imaging. *Radiology* 2010;254(2):384-92. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Grosse C, Globits S, Hergan K. Forty-slice spiral computed tomography of the coronary arteries: assessment of image quality and diagnostic accuracy in a non-selected patient population. *Acta Radiol* 2007;48(1):36-44. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Grossman GB, Garcia EV, Bateman TM, et al. Quantitative Tc-99m sestamibi attenuation-corrected SPECT: development and multicenter trial validation of myocardial perfusion stress gender-independent normal database in an obese population. *J Nucl Cardiol* 2004;11(3):263-72. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Groutars RG, Verzijlbergen JF, Tiel-van Buul MM, et al. The accuracy of 1-day dual-isotope myocardial SPECT in a population with high prevalence of coronary artery disease. *Int J Cardiovasc Imaging* 2003;19(3):229-38. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Grover-McKay M, Milne N, Atwood JE, et al. Comparison of thallium-201 single-photon emission computed tomographic scintigraphy with intravenous dipyridamole and arm exercise. *Am Heart J* 1994;127(6):1516-20. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Grover-McKay M, Ratib O, Schwaiger M,, et al. Detection of coronary artery disease with positron emission tomography and rubidium 82. *Am Heart J* 1992;123(3):646-52. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Groves AM, Speechly-Dick ME, Dickson JC,, et al. Cardiac 82Rubidium PET/CT: initial European experience. *Eur J Nucl Med Mol Imaging* 2007;34(12):1965-72. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Groves AM, Speechly-Dick ME, Kayani I,, et al. First experience of combined cardiac PET/64-detector CT angiography with invasive angiographic validation. *Eur J Nucl Med Mol Imaging* 2009;36(12):2027-33. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Grunig E, Mereles D, Benz A,, et al. Contribution of stress echocardiography to clinical decision making in unselected ambulatory patients with known or suspected coronary artery disease. *Int J Cardiol* 2002;84(2-3):179-85. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Gu X, He Y, Li Z,, et al. Relation between the incidence, location, and extent of thoracic aortic atherosclerosis detected by transesophageal echocardiography and the extent of coronary artery disease by angiography. *Am J Cardiol* 2011;107(2):175-8. *Full-text exclusion reason(s): All women in the study are known to have CAD.*

Gudmundsson P, Shahgaldi K, Winter R,, et al. Head to head comparisons of two modalities of perfusion adenosine stress echocardiography with simultaneous SPECT. *Cardiovasc Ultrasound* 2009;7:19. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Gulati GS, Seth S, Kurian S,, et al. Non-invasive diagnosis of coronary artery disease with 16-slice computed tomography. *Natl Med J India* 2005;18(5):236-41. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Gunalp B, Dokumaci B, Uyan C,, et al. Value of dobutamine technetium-99m-sestamibi SPECT and echocardiography in the detection of coronary artery disease compared with coronary angiography. *J Nucl Med* 1993;34(6):889-94. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Gunes Y, Gumrukcuoglu HA, Kaya Y,, et al. Incremental diagnostic value of color M-mode propagation velocity of the descending thoracic aorta to exercise electrocardiography. *Turk Kardiyoloji Dernegi Arsivi* 2010;38(8):551-557. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Guo SZ, Shu XH, Pan CZ,, et al. Usefulness of dobutamine stress myocardial contrast echocardiography for assessing coronary artery disease. *Chin Med J (Engl)* 2005;118(21):1766-72. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup; No outcomes of interest.*

Gupta NC, Esterbrooks DJ, Hilleman DE,, et al. Comparison of adenosine and exercise thallium-201 single-photon emission computed tomography (SPECT) myocardial perfusion imaging. The GE SPECT Multicenter Adenosine Study Group. *J Am Coll Cardiol* 1992;19(2):248-57. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Haberl R, Becker A, Leber A,, et al. Correlation of coronary calcification and angiographically documented stenoses in patients with suspected coronary artery disease: results of 1,764 patients. *J Am Coll Cardiol* 2001;37(2):451-7. *Full-text exclusion reason(s): No outcomes of interest.*

Haberl R, Tittus J, Bohme E,, et al. Multislice spiral computed tomographic angiography of coronary arteries in patients with suspected coronary artery disease: an effective filter before catheter angiography? *Am Heart J* 2005;149(6):1112-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Hachamovitch R, Berman DS, Shaw LJ,, et al. Incremental prognostic value of myocardial perfusion single photon emission computed tomography for the prediction of cardiac death: differential stratification for risk of cardiac death and myocardial infarction. *Circulation* 1998;97(6):535-43. *Full-text exclusion reason(s): Data for women not reported as a subgroup*

Hachamovitch R, Hayes SW, Friedman JD,, et al. Stress myocardial perfusion single-photon emission computed tomography is clinically effective and cost effective in risk stratification of patients with a high likelihood of coronary artery disease (CAD) but no known CAD. *J Am Coll Cardiol* 2004;43(2):200-8. *Full-text exclusion*

*reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Hachamovitch R, Johnson JR, Hlatky MA, et al. The study of myocardial perfusion and coronary anatomy imaging roles in CAD (SPARC): design, rationale, and baseline patient characteristics of a prospective, multicenter observational registry comparing PET, SPECT, and CTA for resource utilization and clinical outcomes. *J Nucl Cardiol* 2009;16(6):935-48. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Hacker M, Jakobs T, Hack N, et al. Sixty-four slice spiral CT angiography does not predict the functional relevance of coronary artery stenoses in patients with stable angina. *Eur J Nucl Med Mol Imaging* 2007;34(1):4-10. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Hacker M, Jakobs T, Hack N, et al. Combined use of 64-slice computed tomography angiography and gated myocardial perfusion SPECT for the detection of functionally relevant coronary artery stenoses. First results in a clinical setting concerning patients with stable angina. *Nuklearmedizin* 2007;46(1):29-35. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Hacker M, Jakobs T, Matthiesen F, et al. Comparison of spiral multidetector CT angiography and myocardial perfusion imaging in the noninvasive detection of functionally relevant coronary artery lesions: first clinical experiences. *J Nucl Med* 2005;46(8):1294-300. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Hadamitzky M, Distler R, Meyer T, et al. Prognostic value of coronary computed tomographic angiography in comparison with calcium scoring and clinical risk scores. *Circ Cardiovasc Imaging* 2011;4(1):16-23. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Hage FG, Dubovsky EV, Heo J, et al. Outcome of patients with adenosine-induced ST-segment depression but with normal perfusion on tomographic imaging. *Am J Cardiol* 2006;98(8):1009-11. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Hajduczki I, Berenyi I, Enghoff E, et al. Qualitative and quantitative evaluation of the exercise electrocardiogram in assessing the degree of coronary heart disease. *J Electrocardiol* 1985;18(1):55-62. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Hajjiri MM, Leavitt MB, Zheng H, et al. Comparison of positron emission tomography measurement of adenosine-stimulated absolute myocardial blood flow versus relative myocardial tracer content for physiological assessment of coronary artery stenosis severity and location. *JACC Cardiovasc Imaging* 2009;2(6):751-8. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup; No outcomes of interest.*

Halon DA, Gaspar T, Adawi S, et al. Uses and limitations of 40 slice multi-detector row spiral computed tomography for diagnosing coronary lesions in unselected patients referred for routine invasive coronary angiography. *Cardiology* 2007;108(3):200-9. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Halpern EJ, Halpern DJ. Diagnosis of coronary stenosis with CT angiography comparison of automated computer diagnosis with expert readings. *Acad Radiol* 2011;18(3):324-33. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Haluska B, Case C, Short L, et al. Effect of power Doppler and digital subtraction techniques on the comparison of myocardial contrast echocardiography with SPECT. *Heart* 2001;85(5):549-55. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Hambye AS, Vervaet A, Lieber S, et al. Diagnostic value and incremental contribution of bicycle exercise, first-pass radionuclide angiography, and <sup>99m</sup>Tc-labeled sestamibi single-photon emission computed tomography in the identification of coronary artery disease in patients without infarction. *J Nucl Cardiol* 1996;3(6 Pt 1):464-74. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*



Hamdan A, Asbach P, Wellnhofer E,, et al. A prospective study for comparison of MR and CT imaging for detection of coronary artery stenosis. *JACC Cardiovasc Imaging* 2011;4(1):50-61. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup; No outcomes of interest.*

Hamirani YS, Isma'eel H, Larijani V,, et al. The diagnostic accuracy of 64-detector cardiac computed tomography compared with stress nuclear imaging in patients undergoing invasive cardiac catheterization. *J Comput Assist Tomogr* 2010;34(5):645-51. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Hamon M, Fau G, Nee G,, et al. Meta-analysis of the diagnostic performance of stress perfusion cardiovascular magnetic resonance for detection of coronary artery disease. *J Cardiovasc Magn Reson* 2010;12(1):29. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Hamon M, Morello R, Riddell JW. Coronary arteries: diagnostic performance of 16- versus 64-section spiral CT compared with invasive coronary angiography—meta-analysis. *Radiology* 2007;245(3):720-31. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Han PP, Tian YQ, Fang W,, et al. Impact of myocardial perfusion imaging on in-hospital coronary angiography and revascularization of patients with suspected coronary artery disease. *Chin Med J (Engl)* 2011;124(11):1603-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Han SC, Fang CC, Chen Y,, et al. Coronary computed tomography angiography—a promising imaging modality in diagnosing coronary artery disease. *J Chin Med Assoc* 2008;71(5):241-6. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Hansen M, Ginns J, Seneviratne S,, et al. The value of dual-source 64-slice CT coronary angiography in the assessment of patients presenting to an acute chest pain service. *Heart Lung and Circulation* 2010;19(4):213-218. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Hansen PR, Jacobsen TN. Contrast echocardiography as a new non-invasive diagnostic method for assessment of coronary artery disease: from endocardial border detection (EBD) to myocardial perfusion (MCE). *Ugeskrift for laeger* 1998;160(36):5209-5210. *Full-text exclusion reason(s): Non-English.*

Haramati LB, Levsky JM, Jain VR,, et al. CT angiography for evaluation of coronary artery disease in inner-city outpatients: an initial prospective comparison with stress myocardial perfusion imaging. *Int J Cardiovasc Imaging* 2009;25(3):303-13. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Haraphongse M, Kappagoda T, Tymchak W,, et al. The value of sum of ST segment depression in 12-lead electrocardiogram in relation to change in heart rate during exercise to predict the extent of coronary artery disease. *Can J Cardiol* 1986;2(2):64-7. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Harinstein ME, Flaherty JD, Ansari AH,, et al. Predictive value of dobutamine stress echocardiography for coronary artery disease detection in liver transplant candidates. *American Journal of Transplantation* 2008;8(7):1523-1528. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Harshad K, Sathyamurthy I, Ashish G,, et al. Myocardial perfusion single photon emission computed tomography in asymptomatic diabetics. *Indian Heart J* 2010;62(1):29-34. *Full-text exclusion reason(s): Full-text unobtainable.*

Hart CY, Miller TD, Hodge DO,, et al. Specificity of the stress electrocardiogram during adenosine myocardial perfusion imaging in patients taking digoxin. *Am Heart J* 2000;140(6):937-40. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Hatanaka K, Doi M, Hirohata S,, et al. Safety of and tolerance to adenosine infusion for myocardial perfusion single-photon emission computed tomography in a Japanese population. *Circ J* 2007;71(6):904-10. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Hausleiter J, Martinoff S, Hadamitzky M, et al. Image quality and radiation exposure with a low tube voltage protocol for coronary CT angiography results of the PROTECTION II Trial. *JACC Cardiovasc Imaging* 2010;3(11):1113-23. *Full-text exclusion reason(s): Data for women not reported as a subgroup; No outcomes of interest.*

Hausleiter J, Meyer T, Hadamitzky M, et al. Non-invasive coronary computed tomographic angiography for patients with suspected coronary artery disease: the Coronary Angiography by Computed Tomography with the Use of a Submillimeter resolution (CACTUS) trial. *Eur Heart J* 2007;28(24):3034-41. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Hausleiter J, Meyer T, Hermann F, et al. Estimated radiation dose associated with cardiac CT angiography. *JAMA* 2009;301(5):500-7. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Hayashi M, Kambara H, Nohara R, et al. Evaluation and clinical use of early washout ratio of thallium-201 in normal and ischemic myocardium after dipyridamole-induced vasodilation using ring-type emission computed tomography. *Jpn Circ J* 1991;55(3):194-203. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Hayashino Y, Nagata-Kobayashi S, Morimoto T, et al. Cost-effectiveness of screening for coronary artery disease in asymptomatic patients with Type 2 diabetes and additional atherogenic risk factors. *J Gen Intern Med* 2004;19(12):1181-91. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

He Q, Yao Z, Yu X, et al. Evaluation of (99m)Tc-MIBI myocardial perfusion imaging with intravenous infusion of adenosine triphosphate in diagnosis of coronary artery disease. *Chin Med J (Engl)* 2002;115(11):1603-7. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

He ZX, Shi RF, Wu YJ, et al. Direct imaging of exercise-induced myocardial ischemia with fluorine-18-labeled deoxyglucose and Tc-99m-sestamibi in coronary artery disease. *Circulation* 2003;108(10):1208-13. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Hecht HS, Blumfield DE, Hopkins JM, et al. Single dose exercise and redistribution 201thallium scanning in the diagnosis of myocardial ischemia and coronary artery disease. Comparison with exercise and rest electrocardiography, coronary arteriography and left ventriculography. *Chest* 1980;77(3):359-68. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Hecht HS, DeBord L, Shaw R, et al. Supine bicycle stress echocardiography versus tomographic thallium-201 exercise imaging for the detection of coronary artery disease. *J Am Soc Echocardiogr* 1993;6(2):177-85. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Heiba SI, Hayat NJ, Salman HS, et al. Technetium-99m-MIBI myocardial SPECT: supine versus right lateral imaging and comparison with coronary arteriography. *J Nucl Med* 1997;38(10):1510-4. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Heijenbrok-Kal MH, Fleischmann KE, Hunink MG. Stress echocardiography, stress single-photon-emission computed tomography and electron beam computed tomography for the assessment of coronary artery disease: a meta-analysis of diagnostic performance. *Am Heart J* 2007;154(3):415-23. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Heinicke N, Benesch B, Kaiser T, et al. Mechanisms of regional wall motion abnormalities in contrast-enhanced dobutamine stress echocardiography. *Clin Res Cardiol* 2006;95(12):650-6. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Heinle SK, Noblin J, Goree-Best P, et al. Assessment of myocardial perfusion by harmonic power Doppler imaging at rest and during adenosine stress: comparison with (99m)Tc-sestamibi SPECT imaging. *Circulation* 2000;102(1):55-60. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not*

*reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Hendel RC, Layden JJ, Leppo JA. Prognostic value of dipyridamole thallium scintigraphy for evaluation of ischemic heart disease. *J Am Coll Cardiol* 1990;15(1):109-16. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Hennessy TG, Codd MB, Hennessy MS, et al. Comparison of dobutamine stress echocardiography and treadmill exercise electrocardiography for detection of coronary artery disease. *Coron Artery Dis* 1997;8(11-12):689-95. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Hennessy TG, Codd MB, Kane G, et al. Dobutamine stress echocardiography in the detection of coronary artery disease: importance of the pretest likelihood of disease. *Am Heart J* 1997;134(4):685-92. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Hennessy TG, Codd MB, Kane G, et al. Evaluation of patients with diabetes mellitus for coronary artery disease using dobutamine stress echocardiography. *Coron Artery Dis* 1997;8(3-4):171-4. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Hennessy TG, Codd MB, McCarthy C, et al. Dobutamine stress echocardiography in the detection of coronary artery disease in a clinical practice setting. *Int J Cardiol* 1997;62(1):55-62. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Hennessy TG, Siobhan Hennessy M, Codd MB, et al. Detection of coronary artery disease using dobutamine stress echocardiography in patients with an abnormal resting electrocardiograph. *Int J Cardiol* 1998;64(3):293-8. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Henrikson CA, Howell EE, Bush DE, et al. Female sex: A protective role in suspected myocardial ischemia. *Coron Artery Dis* 2006;17(2):153-158. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization.*

Hernandez R, Vale L. The value of myocardial perfusion scintigraphy in the diagnosis and management of angina and myocardial infarction: a probabilistic economic analysis. *Med Decis Making* 2007;27(6):772-88. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Herzog BA, Husmann L, Burkhard N, et al. Accuracy of low-dose computed tomography coronary angiography using prospective electrocardiogram-triggering: first clinical experience. *Eur Heart J* 2008;29(24):3037-42. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Herzog BA, Wyss CA, Husmann L, et al. First head-to-head comparison of effective radiation dose from low-dose 64-slice CT with prospective ECG-triggering versus invasive coronary angiography. *Heart* 2009;95(20):1656-61. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Herzog C, Ay M, Engelmann K, et al. [Visualization techniques in multislice CT-coronary angiography of the heart. Correlations of axial, multiplanar, three-dimensional and virtual endoscopic imaging with the invasive diagnosis]. *Rofo* 2001;173(4):341-9. *Full-text exclusion reason(s): Non-English.*

Herzog C, Britten M, Balzer JO, et al. Multidetector-row cardiac CT: diagnostic value of calcium scoring and CT coronary angiography in patients with symptomatic, but atypical, chest pain. *Eur Radiol* 2004;14(2):169-77. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Herzog C, Nguyen SA, Savino G, et al. Does two-segment image reconstruction at 64-section CT coronary angiography improve image quality and diagnostic accuracy? *Radiology* 2007;244(1):121-9. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Herzog C, Zwerner PL, Doll JR, et al. Significant coronary artery stenosis: comparison on per-patient and per-vessel or per-segment basis at 64-section CT angiography. *Radiology* 2007;244(1):112-20. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Herzog CA, Marwick TH, Pheley AM, et al. Dobutamine stress echocardiography for the detection of significant coronary artery disease in renal transplant candidates. *Am J Kidney Dis* 1999;33(6):1080-90. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Heuschmid M, Burgstahler C, Reimann A, et al. Usefulness of noninvasive cardiac imaging using dual-source computed tomography in an unselected population with high prevalence of coronary artery disease. *Am J Cardiol* 2007;100(4):587-92. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Heuschmid M, Kuettner A, Schroeder S, et al. ECG-gated 16-MDCT of the coronary arteries: assessment of image quality and accuracy in detecting stenoses. *AJR Am J Roentgenol* 2005;184(5):1413-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Hida S, Chikamori T, Tanaka H, et al. Diagnostic value of left ventricular function after adenosine triphosphate loading and at rest in the detection of multi-vessel coronary artery disease using myocardial perfusion imaging. *J Nucl Cardiol* 2009;16(1):20-7. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Hida S, Chikamori T, Tanaka H, et al. Sex-specific approach to gated SPECT volumetric analysis after stress and at rest to detect high-risk coronary artery disease. *Nucl Med Commun* 2010;31(9):800-6. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup; No outcomes of interest.*

Hida S, Chikamori T, Tanaka H, et al. Diagnostic value of left ventricular function after stress and at rest in the detection of multivessel coronary artery disease as assessed by electrocardiogram-gated SPECT. *J Nucl Cardiol* 2007;14(1):68-74. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Higashiue S, Watanabe H, Yokoi Y, et al. Simple detection of severe coronary stenosis using transthoracic Doppler echocardiography at rest. *Am J Cardiol* 2001;87(9):1064-8. *Full-text exclusion reason(s): No data for NITs of interest (ECG, ECHO, SPECT, PET, CMR, CTA).*

Hirai N, Horiguchi J, Fujioka C, et al. Prospective versus retrospective ECG-gated 64-detector coronary CT angiography: assessment of image quality, stenosis, and radiation dose. *Radiology* 2008;248(2):424-30. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Hirano Y, Ozasa Y, Yamamoto T, et al. Diagnosis of vasospastic angina by hyperventilation and cold-pressor stress echocardiography: comparison to I-MIBG myocardial scintigraphy. *J Am Soc Echocardiogr* 2002;15(6):617-23. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Hirano Y, Ozasa Y, Yamamoto T, et al. Hyperventilation and cold-pressor stress echocardiography for noninvasive diagnosis of coronary artery spasm. *J Am Soc Echocardiogr* 2001;14(6):626-33. *Full-text exclusion reason(s): No data for NITs of interest (ECG, ECHO, SPECT, PET, CMR, CTA).*

Hirano Y, Uehara H, Nakamura H, et al. Efficacy of ultrasound-assisted stress testing using a hand-carried ultrasound device for diagnosis of coronary artery disease. *J Am Soc Echocardiogr* 2006;19(5):536-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Hlaihel C, Boussel L, Cochet H, et al. Dose and image quality comparison between prospectively gated axial and retrospectively gated helical coronary CT angiography. *Br J Radiol* 2011;84(997):51-7. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Ho FM, Huang PJ, Liau CS, et al. Dobutamine stress echocardiography compared with dipyridamole thallium-201 single-photon emission computed tomography in detecting coronary artery disease. *Eur Heart J* 1995;16(4):570-5. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Ho JS, Barlow CE, Reinhardt DB, et al. Effect of increasing body mass index on image quality and positive predictive value of 100-kV coronary computed tomographic angiography. *Am J Cardiol* 2010;106(8):1182-1186. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Ho KT, Chua KC, Klotz E,, et al. Stress and rest dynamic myocardial perfusion imaging by evaluation of complete time-attenuation curves with dual-source CT. *JACC: Cardiovascular Imaging* 2010;3(8):811-820. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup; No outcomes of interest.*

Ho YL, Wu CC, Chao CL,, et al. Localizing individual coronary artery obstructions with the dobutamine stress echocardiography. *Cardiology* 1997;88(2):197-202. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Ho YL, Wu CC, Huang PJ,, et al. Dobutamine stress echocardiography compared with exercise thallium-201 single-photon emission computed tomography in detecting coronary artery disease-effect of exercise level on accuracy. *Cardiology* 1997;88(4):379-85. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Ho YL, Wu CC, Lin LC,, et al. Assessment of the coronary artery disease and systolic dysfunction in hypertensive patients with the dobutamine-atropine stress echocardiography: effect of the left ventricular hypertrophy. *Cardiology* 1998;89(1):52-8. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Ho YL, Wu CC, Lin LC,, et al. Assessment of the functional significance of coronary artery stenosis by dobutamine-atropine stress echocardiography. *Cardiology* 1997;88(4):386-92. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Hoekema R, Gürlek C, Brouwer MA,, et al. The use of magnetocardiography and body surface potential mapping in the detection of coronary artery disease in chest pain patients with a normal electrocardiogram. *Computers in Cardiology* 2004;31:389-392. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Hoffmann MH, Shi H, Schmid FT,, et al. Noninvasive coronary imaging with MDCT in comparison to invasive conventional coronary angiography: a fast-developing technology. *AJR Am J Roentgenol* 2004;182(3):601-8. *Full-text exclusion reason(s): Not a clinical study report.*

Hoffmann MH, Shi H, Schmitz BL,, et al. Noninvasive coronary angiography with multislice computed tomography. *JAMA* 2005;293(20):2471-8. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Hoffmann R, Borges AC, Kasprzak JD,, et al. Analysis of myocardial perfusion or myocardial function for detection of regional myocardial abnormalities. An echocardiographic multicenter comparison study using myocardial contrast echocardiography and 2D echocardiography. *Eur J Echocardiogr* 2007;8(6):438-48. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup; No outcomes of interest.*

Hoffmann R, Lethen H, Falter F,, et al. Dobutamine stress echocardiography after coronary artery bypass grafting. Transthoracic vs biplane transoesophageal imaging. *Eur Heart J* 1996;17(2):222-9. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Hoffmann R, Lethen H, Kleinhans E,, et al. Comparative evaluation of bicycle and dobutamine stress echocardiography with perfusion scintigraphy and bicycle electrocardiogram for identification of coronary artery disease. *Am J Cardiol* 1993;72(7):555-9. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup; No outcomes of interest.*

Hoffmann R, Lethen H, Kuhl H,, et al. Extent and severity of test positivity during dobutamine stress echocardiography. Influence on the predictive value for coronary artery disease. *Eur Heart J* 1999;20(20):1485-92. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Hoffmann R, Lethen H, Marwick T,, et al. Analysis of interinstitutional observer agreement in interpretation of dobutamine stress echocardiograms. *J Am Coll Cardiol* 1996;27(2):330-6. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Hoffmann R, Marwick TH, Poldermans D,, et al. Refinements in stress echocardiographic techniques improve inter-institutional agreement in interpretation of dobutamine stress echocardiograms. *Eur Heart J* 2002;23(10):821-9.

*Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Hoffmann S, Mogelvang R, Sogaard P, et al. Tissue Doppler echocardiography reveals impaired cardiac function in patients with reversible ischaemia. *European Journal of Echocardiography* 2011;12(8):628-634. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup*

Hoffmann U, Bamberg F, Chae CU, et al. Coronary computed tomography angiography for early triage of patients with acute chest pain: the ROMICAT (Rule Out Myocardial Infarction using Computer Assisted Tomography) trial. *J Am Coll Cardiol* 2009;53(18):1642-50. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Hoffmann U, Moselewski F, Cury RC, et al. Predictive value of 16-slice multidetector spiral computed tomography to detect significant obstructive coronary artery disease in patients at high risk for coronary artery disease: patient-versus segment-based analysis. *Circulation* 2004;110(17):2638-43. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Hoffmann U, Nagurney JT, Moselewski F, et al. Coronary multidetector computed tomography in the assessment of patients with acute chest pain. *Circulation* 2006;114(21):2251-60. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Hoiland-Carlson PF, Johansen A, Christensen HW, et al. Potential impact of myocardial perfusion scintigraphy as gatekeeper for invasive examination and treatment in patients with stable angina pectoris: observational study without post-test referral bias. *Eur Heart J* 2006;27(1):29-34. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Hollander JE, Chang AM, Shofer FS, et al. One-year outcomes following coronary computerized tomographic angiography for evaluation of emergency department patients with potential acute coronary syndrome. *Acad Emerg Med* 2009;16(8):693-8. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Hollander JE, Litt HI, Chase M, et al. Computed tomography coronary angiography for rapid disposition of low-risk emergency department patients with chest pain syndromes. *Acad Emerg Med* 2007;14(2):112-6. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization.*

Holubkov R, Pepine CJ, Rickens C, et al. Electrocardiogram abnormalities predict angiographic coronary artery disease in women with chest pain: results from the NHLBI WISE Study. *Clin Cardiol* 2002;25(12):553-8. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization.*

Hong YJ, Kim SJ, Lee SM, et al. Low-dose coronary computed tomography angiography using prospective ECG-triggering compared to invasive coronary angiography. *Int J Cardiovasc Imaging* 2010. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Hong YJ, Kim SJ, Lee SM, et al. Low-dose coronary computed tomography angiography using prospective ECG-triggering compared to invasive coronary angiography. *Int J Cardiovasc Imaging* 2011;27(3):425-431. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Hoppe H, Spagnuolo S, Froehlich JM, et al. Retrospective analysis of patients for development of nephrogenic systemic fibrosis following conventional angiography using gadolinium-based contrast agents. *Euro Radiol* 2010;20(3):595-603. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Hosoi M, Sato T, Yamagami K, et al. Impact of diabetes on coronary stenosis and coronary artery calcification detected by electron-beam computed tomography in symptomatic patients. *Diabetes Care* 2002;25(4):696-701. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Hou Y, Yue Y, Guo W, et al. Prospectively versus retrospectively ECG-gated 256-slice coronary CT angiography: image quality and radiation dose over expanded heart rates. *Int J Cardiovasc Imaging* 2010. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Hovland A, Staub UH, Bjornstad H,, et al. Gated SPECT offers improved interobserver agreement compared with echocardiography. *Clin Nucl Med* 2010;35(12):927-30. *Full-text exclusion reason(s): Data for women not reported as a subgroup; No outcomes of interest.*

Hozumi T, Yoshida K, Ogata Y,, et al. Noninvasive assessment of significant left anterior descending coronary artery stenosis by coronary flow velocity reserve with transthoracic color Doppler echocardiography. *Circulation* 1998;97(16):1557-62. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Hsu CC, Chen YW, Hao CL,, et al. Comparison of automated 4D-MSPECT and visual analysis for evaluating myocardial perfusion in coronary artery disease. *Kaohsiung J Med Sci* 2008;24(9):445-52. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Htwe Htwe S, Edell SL, Cacciabardo JM. Noninvasive assessment of coronary artery disease in women after menopause: Gated single proton emission computed tomography myocardial perfusion imaging versus stress echocardiography. *Ultrasound Quarterly* 2000;16(3):151-160. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Hu SJ, Liu SX, Katus HA,, et al. The value of contrast dobutamine stress echocardiography on detecting coronary artery disease in overweight and obese patients. *Can J Cardiol* 2007;23(11):885-9. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Huang PJ, Ho YL, Wu CC,, et al. Simultaneous dobutamine stress echocardiography and thallium-201 perfusion imaging for the detection of coronary artery disease. *Cardiology* 1997;88(6):556-62. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Huang PJ, Yen RF, Chieng PU,, et al. Do beta-blockers affect the diagnostic sensitivity of dobutamine stress thallium-201 single photon emission computed tomographic imaging? *J Nucl Cardiol* 1998;5(1):34-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Huber S, Huber M, Dees D,, et al. Usefulness of multislice spiral computed tomography coronary angiography in patients with acute chest pain in the emergency department. *J Cardiovasc Comput Tomogr* 2007;1(1):29-37. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Huikuri HV, Airaksinen KE, Ikaheimo MJ,, et al. Detection of coronary artery disease by dipyridamole thallium tomography in mitral valve stenosis. *Am J Cardiol* 1989;63(1):124-6. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup.*

Huixing D, Aiqun M, Aimin Y,, et al. Clinical evaluation of Technetium-99m sestamibi myocardial perfusion SPECT in diabetes with suspected coronary artery diseases. *Journal of Medical Colleges of PLA* 2010;25(4):220-225. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Hung GU, Lee KW, Chen CP,, et al. Relationship of transient ischemic dilation in dipyridamole myocardial perfusion imaging and stress-induced changes of functional parameters evaluated by TI-201 gated SPECT. *J Nucl Cardiol* 2005;12(3):268-75. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Hung GU, Lee KW, Chen CP,, et al. Worsening of left ventricular ejection fraction induced by dipyridamole on TI-201 gated myocardial perfusion imaging predicts significant coronary artery disease. *J Nucl Cardiol* 2006;13(2):225-32. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Hur J, Kim YJ, Lee HJ,, et al. Quantification and characterization of obstructive coronary plaques using 64-slice computed tomography: a comparison with intravascular ultrasound. *J Comput Assist Tomogr* 2009;33(2):186-92. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Husmann L, Gaemperli O, Schepis T,, et al. Accuracy of quantitative coronary angiography with computed tomography and its dependency on plaque composition: plaque composition and accuracy of cardiac CT. *Int J Cardiovasc Imaging* 2008;24(8):895-904. *Full-text exclusion reason(s): No women with symptomatic chest pain, or*

*results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Husmann L, Herzog BA, Burger IA, et al. Usefulness of additional coronary calcium scoring in low-dose CT coronary angiography with prospective ECG-triggering impact on total effective radiation dose and diagnostic accuracy. *Acad Radiol* 2010;17(2):201-6. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Husmann L, Herzog BA, Gaemperli O, et al. Diagnostic accuracy of computed tomography coronary angiography and evaluation of stress-only single-photon emission computed tomography/computed tomography hybrid imaging: comparison of prospective electrocardiogram-triggering vs. retrospective gating. *Eur Heart J* 2009;30(5):600-7. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Husmann L, Scheffel H, Valenta I, et al. Impact of hypertension on the diagnostic accuracy of coronary angiography with computed tomography. *Int J Cardiovasc Imaging* 2008;24(7):763-70. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Husmann L, Schepis T, Scheffel H, et al. Comparison of diagnostic accuracy of 64-slice computed tomography coronary angiography in patients with low, intermediate, and high cardiovascular risk. *Acad Radiol* 2008;15(4):452-61. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Husmann L, Valenta I, Gaemperli O, et al. Feasibility of low-dose coronary CT angiography: first experience with prospective ECG-gating. *Eur Heart J* 2008;29(2):191-7. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Husmann L, Wiegand M, Valenta I, et al. Diagnostic accuracy of myocardial perfusion imaging with single photon emission computed tomography and positron emission tomography: a comparison with coronary angiography. *Int J Cardiovasc Imaging* 2008;24(5):511-8. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Husser O, Bodi V, Sanchis J, et al. Additional diagnostic value of systolic dysfunction induced by dipyridamole stress cardiac magnetic resonance used in detecting coronary artery disease. *Rev Esp Cardiol* 2009;62(4):383-91. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Hwang Y, Kim Y, Chung IM, et al. Coronary heart disease risk assessment and characterization of coronary artery disease using coronary CT angiography: comparison of asymptomatic and symptomatic groups. *Clin Radiol* 2010;65(8):601-8. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; No outcomes of interest.*

Iglesias-Garriz I, Rodriguez MA, Garcia-Porrero E, et al. Emergency nontraumatic chest pain: use of stress echocardiography to detect significant coronary artery stenosis. *J Am Soc Echocardiogr* 2005;18(11):1181-6. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Ikonen AE, Manninen HI, Vainio P, et al. Three-dimensional respiratory-gated coronary MR angiography with reference to X-ray coronary angiography. *Acta Radiol* 2003;44(6):583-9. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Imamura Y, Fukuyama T, Nishimura S, et al. Normal myocardial perfusion scan portends a benign prognosis independent from the pretest probability of coronary artery disease. Sub-analysis of the J-ACCESS study. *J Cardiol* 2009;54(1):93-100. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*



Ingul CB, Stoylen A, Slordahl SA, et al. Automated analysis of myocardial deformation at dobutamine stress echocardiography: an angiographic validation. *J Am Coll Cardiol* 2007;49(15):1651-9. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Isaac D, Walling A. Clinical evaluation of women with ischemic heart disease: diagnosis and noninvasive testing. *Can J Cardiol* 2001;17 Suppl D:38D-48D. *Full-text exclusion reason(s): Not a clinical study report.*

Ishida N, Sakuma H, Motoyasu M, et al. Noninfarcted myocardium: correlation between dynamic first-pass contrast-enhanced myocardial MR imaging and quantitative coronary angiography. *Radiology* 2003;229(1):209-16. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Ishii K, Imai M, Suyama T, et al. Exercise-induced post-ischemic left ventricular delayed relaxation or diastolic stunning: is it a reliable marker in detecting coronary artery disease? *J Am Coll Cardiol* 2009;53(8):698-705. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Iskandrian AE, Heo J, Nallamothu N. Detection of coronary artery disease in women with use of stress single-photon emission computed tomography myocardial perfusion imaging. *J Nucl Cardiol* 1997;4(4):329-35. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Iskandrian AS, Chae SC, Heo J, et al. Independent and incremental prognostic value of exercise single-photon emission computed tomographic (SPECT) thallium imaging in coronary artery disease. *J Am Coll Cardiol* 1993;22(3):665-70. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Iskandrian AS, Hakki AH, Kane-Marsch S. Exercise thallium-201 scintigraphy in men with nondiagnostic exercise electrocardiograms. Prognostic implications. *Arch Intern Med* 1986;146(11):2189-93. *Full-text exclusion reason(s): Population does not include women  $\geq$  age 18; Data for women not reported as a subgroup.*

Iskandrian AS, Hakki AH, Kane-Marsch S. Prognostic implications of exercise thallium-201 scintigraphy in patients with suspected or known coronary artery disease. *Am Heart J* 1985;110(1 Pt 1):135-43. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Iskandrian AS, Heo J, Decoskey D, et al. Use of exercise thallium-201 imaging for risk stratification of elderly patients with coronary artery disease. *Am J Cardiol* 1988;61(4):269-72. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Iskandrian AS, Heo J, Kong B, et al. Effect of exercise level on the ability of thallium-201 tomographic imaging in detecting coronary artery disease: analysis of 461 patients. *J Am Coll Cardiol* 1989;14(6):1477-86. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Iskandrian AS, Johnson J, Le TT, et al. Comparison of the treadmill exercise score and single-photon emission computed tomographic thallium imaging in risk assessment. *J Nucl Cardiol* 1994;1(2 Pt 1):144-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Isma'eel H, Shamseddeen W, El Khoury M, et al. Diabetes supersedes dobutamine stress echocardiography in predicting cardiac events in female patients. *Clin Exp Obstet Gynecol* 2010;37(3):197-200. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization.*

Ismail G, Lo E, Sada M, et al. Long-term prognosis of patients with a normal exercise echocardiogram and clinical suspicion of myocardial ischemia. *Am J Cardiol* 1995;75(14):934-6. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Isobe S, Sato K, Sugiura K, et al. Feasibility of intravenous administration of landiolol hydrochloride for multislice computed tomography coronary angiography: initial experience. *Circ J* 2008;72(11):1814-20. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Iwasaki K, Matsumoto T, Aono H, et al. Prevalence of subclinical atherosclerosis in asymptomatic patients with low-to-intermediate risk by 64-slice computed tomography. *Coron Artery Dis* 2011;22(1):18-25. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Iwasaki K, Matsumoto T, Aono H, et al. Relationship between epicardial fat measured by 64-multidetector computed tomography and coronary artery disease. *Clin Cardiol* 2011;34(3):166-71. *Full-text exclusion reason(s): Data for women not reported as a subgroup; No outcomes of interest.*

Iwase M, Fukui M, Tamagaki H, et al. Advantages and disadvantages of dobutamine stress echocardiography compared with treadmill exercise electrocardiography in detecting ischemia. *Jpn Circ J* 1996;60(12):954-60. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Jahnke C, Gebker R, Manka R, et al. Navigator-gated 3D blood oxygen level-dependent CMR at 3.0-T for detection of stress-induced myocardial ischemic reactions. *JACC Cardiovasc Imaging* 2010;3(4):375-84. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Jahnke C, Paetsch I, Nehrke K, et al. Rapid and complete coronary arterial tree visualization with magnetic resonance imaging: feasibility and diagnostic performance. *Eur Heart J* 2005;26(21):2313-9. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Jahnke C, Paetsch I, Schnackenburg B, et al. Coronary MR angiography with steady-state free precession: individually adapted breath-hold technique versus free-breathing technique. *Radiology* 2004;232(3):669-76. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Jahnke C, Paetsch I, Schnackenburg B, et al. Comparison of radial and Cartesian imaging techniques for MR coronary angiography. *J Cardiovasc Magn Reson* 2004;6(4):865-75. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Jain A, McClelland RL, Polak JF, et al. Cardiovascular imaging for assessing cardiovascular risk in asymptomatic men versus women: the multi-ethnic study of atherosclerosis (MESA). *Circ Cardiovasc Imaging* 2011;4(1):8-15. *Full-text exclusion reason(s): No data for NITs of interest (ECG, ECHO, SPECT, PET, CMR, CTA).*

Janne d'Othee B, Siebert U, Cury R, et al. A systematic review on diagnostic accuracy of CT-based detection of significant coronary artery disease. *Eur J Radiol* 2008;65(3):449-61. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Javadrashid R, Salehi A, Tarzamni MK, et al. Diagnostic efficacy of coronary calcium score in the assessment of significant coronary artery stenosis. *Kardiol Pol* 2010;68(3):285-91. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Jeetley P, Burden L, Greaves K, et al. Prognostic value of myocardial contrast echocardiography in patients presenting to hospital with acute chest pain and negative troponin. *Am J Cardiol* 2007;99(10):1369-73. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Jeetley P, Burden L, Senior R. Stress echocardiography is superior to exercise ECG in the risk stratification of patients presenting with acute chest pain with negative Troponin. *Eur J Echocardiogr* 2006;7(2):155-64. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Jeetley P, Burden L, Stoykova B, et al. Clinical and economic impact of stress echocardiography compared with exercise electrocardiography in patients with suspected acute coronary syndrome but negative troponin: a prospective randomized controlled study. *Eur Heart J* 2007;28(2):204-11. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Jeetley P, Hickman M, Kamp O, et al. Myocardial contrast echocardiography for the detection of coronary artery stenosis: a prospective multicenter study in comparison with single-photon emission computed tomography. *J Am Coll Cardiol* 2006;47(1):141-5. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Jelinek JS, Murphey MD, Welker JA, et al. Diagnosis of primary bone tumors with image-guided percutaneous biopsy: experience with 110 tumors. *Radiology* 2002;223(3):731-7. *Full-text exclusion reason(s): No data for NITs of interest (ECG, ECHO, SPECT, PET, CMR, CTA).*

Jengo JA, Freeman R, Brizendine M, et al. Detection of coronary artery disease: comparison of exercise stress radionuclide angiography and thallium stress perfusion scanning. *Am J Cardiol* 1980;45(3):535-41. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Jeong DW, Choo KS, Baik SK, et al. Step-and-shoot prospectively ECG-gated versus retrospectively ECG-gated with tube current modulation coronary CT angiography using the 128-slice MDCT: comparison of image quality and radiation dose. *Acta Radiol* 2011;52(2):155-60. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Jeong HC, Ahn Y, Ko JS, et al. The role of 64-slice multi-detector computed tomography in the detection of subclinical atherosclerosis of the coronary artery. *Int J Cardiovasc Imaging* 2010;26(Suppl 2):253-9. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Jiang B, Gai L, Sun Z, et al. The combination of 64 multislice CT angiography and optical coherence tomography optimally characterizes coronary plaques. *Acta Cardiol* 2011;66(2):213-8. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup; No outcomes of interest.*

Jimenez-Hoyuela Garcia JM, Robledo Carmona J, Ortega Lozano S, et al. Myocardial perfusion scintigraphy in the emergency department for the evaluation and triage of patients with chest pain and a non-diagnostic electrocardiogram [Gammagrafia de perfusion miocardica en el servicio de urgencias para la evaluacion y triage de pacientes con dolor toracico y electrocardiograma no diagnostico]. *Investigacion Cardiovascular* 2006;9(1):7-18. *Full-text exclusion reason(s): Full-text unobtainable.*

Jinzaki M, Sato K, Tanami Y, et al. Diagnostic accuracy of angiographic view image for the detection of coronary artery stenoses by 64-detector row CT: a pilot study comparison with conventional post-processing methods and axial images alone. *Circ J* 2009;73(4):691-8. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Johansen AH, Poulsen TS, Hoiland-Carlsen PF, et al. Myocardial perfusion imaging and coronary angiography in patients with known or suspected stable angina pectoris. *Dan Med Bull* 2001;48(2):80-3. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

John RM, Taggart PI, Sutton PM, et al. Vasodilator myocardial perfusion imaging: demonstration of local electrophysiological changes of ischaemia. *Br Heart J* 1992;68(1):21-30. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Johnson NP, Schimmel DR, Jr., Dyer SP, et al. Survival by stress modality in patients with a normal myocardial perfusion study. *Am J Cardiol* 2011;107(7):986-9. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Johnson TR, Nikolaou K, Busch S, et al. Diagnostic accuracy of dual-source computed tomography in the diagnosis of coronary artery disease. *Invest Radiol* 2007;42(10):684-91. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Johnson TR, Nikolaou K, Wintersperger BJ, et al. ECG-gated 64-MDCT angiography in the differential diagnosis of acute chest pain. *AJR Am J Roentgenol* 2007;188(1):76-82. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Joint Commission. Radiation risks of diagnostic imaging. *Sentinel Event Alert* 2011(47):1-4. *Full-text exclusion reason(s): Not a clinical study report.*

Jonsbu E, Dammen T, Morken G, et al. Cardiac and psychiatric diagnoses among patients referred for chest pain and palpitations. *Scand Cardiovasc J* 2009;43(4):256-9. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup; No outcomes of interest.*

Kachenoura N, Gaspar T, Lodato JA, et al. Combined assessment of coronary anatomy and myocardial perfusion using multidetector computed tomography for the evaluation of coronary artery disease. *Am J Cardiol* 2009;103(11):1487-94. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Kaiser C, Bremerich J, Haller S,, et al. Limited diagnostic yield of non-invasive coronary angiography by 16-slice multi-detector spiral computed tomography in routine patients referred for evaluation of coronary artery disease. *Eur Heart J* 2005;26(19):1987-92. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Kajander S, Joutsiniemi E, Saraste M,, et al. Cardiac positron emission tomography/computed tomography imaging accurately detects anatomically and functionally significant coronary artery disease. *Circulation* 2010;122(6):603-13. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Kajinami K, Seki H, Takekoshi N,, et al. Noninvasive prediction of coronary atherosclerosis by quantification of coronary artery calcification using electron beam computed tomography: comparison with electrocardiographic and thallium exercise stress test results. *J Am Coll Cardiol* 1995;26(5):1209-21. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup.*

Kaltenbach M, Martin KL, Hopf R. [Accuracy of exercise tests in the recognition of coronary-artery stenosis. Comparison between post-exercise ECG and coronary arteriogram (author's transl)]. *Dtsch Med Wochenschr* 1976;101(52):1907-11. *Full-text exclusion reason(s): Non-English.*

Kamal AM, Fattah AA, Pancholy S,, et al. Prognostic value of adenosine single-photon emission computed tomographic thallium imaging in medically treated patients with angiographic evidence of coronary artery disease. *J Nucl Cardiol* 1994;1(3):254-61. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Kaminek M, Myslivecek M, Skvarilova M,, et al. [Prognostic significance of stress tomographic scintigraphy of myocardial perfusion in diabetic patients]. *Vnitr Lek* 2001;47(11):739-43. *Full-text exclusion reason(s): Non-English.*

Kaneko K, Ito M, Takanashi T,, et al. Computed tomography and scintigraphy vs. cardiac catheterization for coronary disease screening prior to noncardiac surgery. *Internal Medicine* 2010;49(16):1703-1710. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Kang DH, Kang SJ, Song JM,, et al. Efficacy of myocardial contrast echocardiography in the diagnosis and risk stratification of acute coronary syndrome. *Am J Cardiol* 2005;96(11):1498-502. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Kang DK, Im NJ, Park SM,, et al. CT comparison of visual and computerised quantification of coronary stenosis according to plaque composition. *Eur Radiol* 2011;21(4):712-21. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Kang HJ, Lee HY, Na SH,, et al. Differential effect of intracoronary infusion of mobilized peripheral blood stem cells by granulocyte colony-stimulating factor on left ventricular function and remodeling in patients with acute myocardial infarction versus old myocardial infarction: the MAGIC Cell-3-DES randomized, controlled trial. *Circulation* 2006;114(1 Suppl):I145-51. *Full-text exclusion reason(s): No data for NITs of interest (ECG, ECHO, SPECT, PET, CMR, CTA).*

Kanzaki H, Nakatani S, Kandori A,, et al. A new screening method to diagnose coronary artery disease using multichannel magnetocardiogram and simple exercise. *Basic Res Cardiol* 2003;98(2):124-32. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Karabinos IK, Papadopoulos A, Karvouni E,, et al. Reliability and safety of dobutamine stress echocardiography for detection of myocardial ischemia-viability: Experience from 802 consecutive studies. *Hellenic Journal of Cardiology* 2004;45(2):71-83. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Karagiannis SE, Bax JJ, Elhendy A,, et al. Enhanced sensitivity of dobutamine stress echocardiography by observing wall motion abnormalities during the recovery phase after acute beta-blocker administration. *Am J Cardiol* 2006;97(4):462-5. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Karamitsos TD, Arnold JR, Pegg TJ,, et al. Tolerance and safety of adenosine stress perfusion cardiovascular magnetic resonance imaging in patients with severe coronary artery disease. *Int J Cardiovasc Imaging*

2009;25(3):277-83. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Kasalicky J, Kovac I, Lanska V. Pretest clinical diagnosis of coronary artery disease and stress myocardial perfusion scintigram. *Nucl Med Rev Cent East Eur* 2001;4(2):89-92. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup; No outcomes of interest.*

Kasprzak JD, Ciesielczyk M, Krzeminska-Pakula M., et al. Myocardial contrast echocardiography using dipyridamole-atropine stress - Preliminary report. *Polski Przegląd Kardiologiczny* 2002;4(2):141-148. *Full-text exclusion reason(s): Non-English.*

Katayama T, Ogata N, Tsuruya Y. Diagnostic accuracy of supine and prone thallium-201 stress myocardial perfusion single-photon emission computed tomography to detect coronary artery disease in inferior wall of left ventricle. *Ann Nucl Med* 2008;22(4):317-21. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Kato S, Kitagawa K, Ishida N., et al. Assessment of coronary artery disease using magnetic resonance coronary angiography: A national multicenter trial. *J Am Coll Cardiol* 2010;56(12):983-991. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Kaul S, Boucher CA, Newell JB., et al. Determination of the quantitative thallium imaging variables that optimize detection of coronary artery disease. *J Am Coll Cardiol* 1986;7(3):527-37. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Kaul S, Finkelstein DM, Homma S., et al. Superiority of quantitative exercise thallium-201 variables in determining long-term prognosis in ambulatory patients with chest pain: a comparison with cardiac catheterization. *J Am Coll Cardiol* 1988;12(1):25-34. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Kaul S, Lilly DR, Gascho JA., et al. Prognostic utility of the exercise thallium-201 test in ambulatory patients with chest pain: comparison with cardiac catheterization. *Circulation* 1988;77(4):745-58. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Kawai Y, Morita K, Nozaki Y., et al. Diagnostic value of (123)I-beta-methyl-p-iodophenyl-pentadecanoic acid (BMIPP) single photon emission computed tomography (SPECT) in patients with chest pain - Comparison with rest-stress (99m)Tc-tetrofosmin SPECT and coronary angiography. *Circ J* 2004;68(6):547-552. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Kawase Y, Nishimoto M, Hato K., et al. Assessment of coronary artery disease with nicorandil stress magnetic resonance imaging. *Osaka City Med J* 2004;50(2):87-94. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Kefer J, Coche E, Legros G., et al. Head-to-head comparison of three-dimensional navigator-gated magnetic resonance imaging and 16-slice computed tomography to detect coronary artery stenosis in patients. *J Am Coll Cardiol* 2005;46(1):92-100. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup; No outcomes of interest.*

Kelle S, Hamdan A, Schnackenburg B., et al. Dobutamine stress cardiovascular magnetic resonance at 3 Tesla. *J Cardiovasc Magn Reson* 2008;10:44. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Ketelsen D, Fenchel M, Thomas C., et al. Estimation of radiation exposure of retrospective gated and prospective triggered 128-slice triple-rule-out CT angiography. *Acta Radiol* 2011. *Full-text exclusion reason(s): Not a clinical study report; Population does not include women  $\geq$  age 18.*

Khan A, Nasir K, Khosa F., et al. Prospective gating with 320-MDCT angiography: effect of volume scan length on radiation dose. *AJR Am J Roentgenol* 2011;196(2):407-11. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Khan MF, Wesarg S, Gurung J, et al. Facilitating coronary artery evaluation in MDCT using a 3D automatic vessel segmentation tool. *Eur Radiol* 2006;16(8):1789-95. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup; No outcomes of interest.*

Khare RK, Courtney DM, Powell ES, et al. Sixty-four-slice computed tomography of the coronary arteries: cost-effectiveness analysis of patients presenting to the emergency department with low-risk chest pain. *Acad Emerg Med* 2008;15(7):623-32. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Khattar RS, Senior R, Lahiri A. Assessment of myocardial perfusion and contractile function by inotropic stress Tc-99m sestamibi SPECT imaging and echocardiography for optimal detection of multivessel coronary artery disease. *Heart* 1998;79(3):274-80. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Khorsand A, Graf S, Sochor H, et al. Automated assessment of myocardial SPECT perfusion scintigraphy: a comparison of different approaches of case-based reasoning. *Artif Intell Med* 2007;40(2):103-13. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup; No outcomes of interest.*

Khorsand A, Haddad M, Graf S, et al. Automated assessment of dipyridamole 201Tl myocardial SPECT perfusion scintigraphy by case-based reasoning. *J Nucl Med* 2001;42(2):189-93. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup; No outcomes of interest.*

Kiat H, Maddahi J, Roy LT, et al. Comparison of technetium 99m methoxy isobutyl isonitrile and thallium 201 for evaluation of coronary artery disease by planar and tomographic methods. *Am Heart J* 1989;117(1):1-11. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Kim C, Kwok YS, Heagerty P, et al. Pharmacologic stress testing for coronary disease diagnosis: A meta-analysis. *Am Heart J* 2001;142(6):934-44. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Kim J, Lee H, Song S, et al. Efficacy and safety of the computed tomography coronary angiography based approach for patients with acute chest pain at an emergency department: one month clinical follow-up study. *J Korean Med Sci* 2010;25(3):466-71. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Kim SY, Kim KS, Seung MJ, et al. The culprit lesion score on multi-detector computed tomography can detect vulnerable coronary artery plaque. *Int J Cardiovasc Imaging* 2010;26(Suppl 2):245-52. *Full-text exclusion reason(s): Data for women not reported as a subgroup; No outcomes of interest.*

Kim TH, Yu SH, Choi SH, et al. Pericardial fat amount is an independent risk factor of coronary artery stenosis assessed by multidetector-row computed tomography: The Korean atherosclerosis study 2. *Obesity* 2011;19(5):1028-1034. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Kim WY, Danias PG, Stuber M, et al. Coronary magnetic resonance angiography for the detection of coronary stenoses. *N Engl J Med* 2001;345(26):1863-9. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Kim YJ, Seo JS, Choi BW, et al. Feasibility and diagnostic accuracy of whole heart coronary MR angiography using free-breathing 3D balanced turbo-field-echo with SENSE and the half-fourier acquisition technique. *Korean J Radiol* 2006;7(4):235-42. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Kirchin MA, Pirovano G, Venetianer C, et al. Safety assessment of gadobenate dimeglumine (MultiHance): extended clinical experience from phase I studies to post-marketing surveillance. *J Magn Reson Imaging* 2001;14(3):281-94. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization.*

Kiriyama T, Toba M, Fukushima Y, et al. Discordance between the morphological and physiological information of 64-slice MSCT coronary angiography and myocardial perfusion imaging in patients with intermediate to high probability of coronary artery disease. *Circ J* 2011;75(7):1670-1677. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Kirschbaum SW, Nieman K, Springeling T, et al. Non-invasive diagnostic workup of patients with suspected stable angina by combined computed tomography coronary angiography and magnetic resonance perfusion imaging. *Circ J* 2011;75(7):1678-1684. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Kisacik HL, Ozdemir K, Altinyay E, et al. Comparison of exercise stress testing with simultaneous dobutamine stress echocardiography and technetium-99m isonitrile single-photon emission computerized tomography for diagnosis of coronary artery disease. *Eur Heart J* 1996;17(1):113-9. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Kitagawa K, Sakuma H, Nagata M, et al. Diagnostic accuracy of stress myocardial perfusion MRI and late gadolinium-enhanced MRI for detecting flow-limiting coronary artery disease: a multicenter study. *Eur Radiol* 2008;18(12):2808-16. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Klein AJ, Garcia JA, Hudson PA, et al. Safety and efficacy of dual-axis rotational coronary angiography vs. standard coronary angiography. *Catheter Cardiovasc Interv* 2011;77(6):820-7. *Full-text exclusion reason(s): No data for NITs of interest (ECG, ECHO, SPECT, PET, CMR, CTA); Data for women not reported as a subgroup.*

Klein C, Gebker R, Kokocinski T, et al. Combined magnetic resonance coronary artery imaging, myocardial perfusion and late gadolinium enhancement in patients with suspected coronary artery disease. *J Cardiovasc Magn Reson* 2008;10:45. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Klem I, Heitner JF, Shah DJ, et al. Improved detection of coronary artery disease by stress perfusion cardiovascular magnetic resonance with the use of delayed enhancement infarction imaging. *J Am Coll Cardiol* 2006;47(8):1630-8. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Klumpp B, Hoevelborn T, Fenchel M, et al. Magnetic resonance myocardial perfusion imaging-First experience at 3.0T. *Eur J Radiol* 2009;69(1):165-72. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Klumpp BD, Seeger A, Doesch C, et al. High resolution myocardial magnetic resonance stress perfusion imaging at 3 T using a 1 M contrast agent. *Eur Radiol* 2010;20(3):533-41. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Knez A, Becker CR, Leber A, et al. Usefulness of multislice spiral computed tomography angiography for determination of coronary artery stenoses. *Am J Cardiol* 2001;88(10):1191-4. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Ko SM, Kim NR, Kim DH, et al. Assessment of image quality and radiation dose in prospective ECG-triggered coronary CT angiography compared with retrospective ECG-gated coronary CT angiography. *Int J Cardiovasc Imaging* 2010;26 Suppl 1:93-101. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Koh AS, Flores JLS, Keng FYJ, et al. Evaluation of the American College of Cardiology Foundation/American Society of Nuclear Cardiology appropriateness criteria for SPECT myocardial perfusion imaging in an Asian tertiary cardiac center. *J Nucl Cardiol* 2011;18(2):323-330. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup; No outcomes of interest.*

Koide Y, Yotsukura M, Yoshino H, et al. Usefulness of QT dispersion immediately after exercise as an indicator of coronary stenosis independent of gender or exercise-induced ST-segment depression. *Am J Cardiol* 2000;86(12):1312-7. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Koide Y, Yotsukura M, Yoshino H, et al. Value of QT dispersion in the interpretation of treadmill exercise electrocardiograms of patients without exercise-induced chest pain or ST-segment depression. *Am J Cardiol* 2000;85(9):1094-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Kolnes K, Velle OH, Hareide S, et al. Multislice computed tomography coronary angiography at a local hospital: Pitfalls and potential. *Acta Radiol* 2006;47(7):680-6. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Kong G, Lichtenstein M, Gunawardana D, et al. Fixed defect on rest/stress Tc-99m sestamibi study underestimates myocardial ischemia: comparison with 24-hour thallium-201 study for short- and intermediate-term follow-up. *Clin Nucl Med* 2008;33(3):168-71. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Konieczynska M, Tracz W, Pasowicz M, et al. Use of coronary calcium score in the assessment of atherosclerotic lesions in coronary arteries. *Kardiol Pol* 2006;64(10):1073-9; discussion 1080-1. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Kontos MC, Dilsizian V, Weiland F, et al. Iodoflitic acid I 123 (BMIPP) fatty acid imaging improves initial diagnosis in emergency department patients with suspected acute coronary syndromes: a multicenter trial. *J Am Coll Cardiol* 2010;56(4):290-9. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Kontos MC, Hinchman D, Cunningham M, et al. Comparison of contrast echocardiography with single-photon emission computed tomographic myocardial perfusion imaging in the evaluation of patients with possible acute coronary syndromes in the emergency department. *Am J Cardiol* 2003;91(9):1099-102. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Kontos MC, Jesse RL, Tatum JL, et al. Coronary angiographic findings in patients with cocaine-associated chest pain. *J Emerg Med* 2003;24(1):9-13. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup; No outcomes of interest.*

Kontos MC, Schmidt KL, McCue M, et al. A comprehensive strategy for the evaluation and triage of the chest pain patient: a cost comparison study. *J Nucl Cardiol* 2003;10(3):284-90. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup; No outcomes of interest.*

Kopp AF, Schroeder S, Kuettner A, et al. Non-invasive coronary angiography with high resolution multidetector-row computed tomography. Results in 102 patients. *Eur Heart J* 2002;23(21):1714-25. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Korosoglou G, da Silva KG, Jr., Labadze N, et al. Real-time myocardial contrast echocardiography for pharmacologic stress testing: is quantitative estimation of myocardial blood flow reserve necessary? *J Am Soc Echocardiogr* 2004;17(1):1-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Korosoglou G, Dubart AE, DaSilva KG, Jr., et al. Real-time myocardial perfusion imaging for pharmacologic stress testing: added value to single photon emission computed tomography. *Am Heart J* 2006;151(1):131-8. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Korosoglou G, Elhmidi Y, Steen H, et al. Prognostic value of high-dose dobutamine stress magnetic resonance imaging in 1,493 consecutive patients: assessment of myocardial wall motion and perfusion. *J Am Coll Cardiol* 2010;56(15):1225-34. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Korosoglou G, Labadze N, Hansen A, et al. Usefulness of real-time myocardial perfusion imaging in the evaluation of patients with first time chest pain. *Am J Cardiol* 2004;94(10):1225-31. *Full-text exclusion reason(s): All women in the study are known to have CAD.*

Korosoglou G, Lehrke S, Wochele A, et al. Strain-encoded CMR for the detection of inducible ischemia during intermediate stress. *JACC Cardiovasc Imaging* 2010;3(4):361-71. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*



Kostkiewicz M, Konieczynska M, Szot WM, et al. Comparison between (99m)Tc-MIBI myocardial perfusion SPECT and multi-slice computed tomography for identifying and assessing coronary artery disease. *Hellenic Journal of Nuclear Medicine* 2004;7(1):48-51. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Kowatsch I, Tsutsui JM, Osorio AF, et al. Head-to-head comparison of dobutamine and adenosine stress real-time myocardial perfusion echocardiography for the detection of coronary artery disease. *J Am Soc Echocardiogr* 2007;20(9):1109-17. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Krenning BJ, Geleijnse ML, Poldermans D, et al. Methodological analysis of diagnostic dobutamine stress echocardiography studies. *Echocardiography* 2004;21(8):725-36. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Krenning BJ, Nemes A, Soliman OI, et al. Contrast-enhanced three-dimensional dobutamine stress echocardiography: between Scylla and Charybdis? *Eur J Echocardiogr* 2008;9(6):757-60. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Krishnam MS, Tomasian A, Iv M, et al. Left ventricular ejection fraction using 64-slice CT coronary angiography and new evaluation software: initial experience. *Br J Radiol* 2008;81(966):450-5. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup; No outcomes of interest.*

Krittayaphong R, Boonyasirinant T, Saiviroonporn P, et al. Myocardial perfusion cardiac magnetic resonance for the diagnosis of coronary artery disease: do we need rest images? *Int J Cardiovasc Imaging* 2009;25 Suppl 1:139-48. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Krittayaphong R, Chaithiraphan V, Maneesai A, et al. Prognostic value of combined magnetic resonance myocardial perfusion imaging and late gadolinium enhancement. *Int J Cardiovasc Imaging* 2011;27(5):705-14. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Krittayaphong R, Mahanonda N, Kangkagate C, et al. Accuracy of magnetic resonance imaging in the diagnosis of coronary artery disease. *J Med Assoc Thai* 2003;86 Suppl 1:S59-66. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Krittayaphong R, Saiviroonporn P, Boonyasirinant T, et al. Prevalence and prognosis of myocardial scar in patients with known or suspected coronary artery disease and normal wall motion. *J Cardiovasc Magn Reson* 2011;13(1):2. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Krzanowski M, Bodzon W, Brzostek T, et al. Value of transthoracic echocardiography for the detection of high-grade coronary artery stenosis: prospective evaluation in 50 consecutive patients scheduled for coronary angiography. *J Am Soc Echocardiogr* 2000;13(12):1091-9. *Full-text exclusion reason(s): No data for NITs of interest (ECG, ECHO, SPECT, PET, CMR, CTA).*

Kuettner A, Beck T, Drosch T, et al. Diagnostic accuracy of noninvasive coronary imaging using 16-detector slice spiral computed tomography with 188 ms temporal resolution. *J Am Coll Cardiol* 2005;45(1):123-7. *Full-text exclusion reason(s): No outcomes of interest.*

Kuettner A, Beck T, Drosch T, et al. Image quality and diagnostic accuracy of non-invasive coronary imaging with 16 detector slice spiral computed tomography with 188 ms temporal resolution. *Heart* 2005;91(7):938-41. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Kuettner A, Trabold T, Schroeder S, et al. Noninvasive detection of coronary lesions using 16-detector multislice spiral computed tomography technology: initial clinical results. *J Am Coll Cardiol* 2004;44(6):1230-7. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Kuijpers D, van Dijkman PR, Janssen CH, et al. Dobutamine stress MRI. Part II. Risk stratification with dobutamine cardiovascular magnetic resonance in patients suspected of myocardial ischemia. *Eur Radiol*

2004;14(11):2046-52. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Kumar R, Patel CD, Marwah A,, et al. Detection of coronary artery disease by stress thallium scintigraphy in diabetic patients. *Nucl Med Commun* 2001;22(3):287-9. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Kunimasa T, Sato Y, Matsumoto N,, et al. Detection of coronary artery disease by free-breathing, whole heart coronary magnetic resonance angiography: our initial experience. *Heart Vessels* 2009;24(6):429-33. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Kuntz KM, Fleischmann KE, Hunink MG,, et al. Cost-effectiveness of diagnostic strategies for patients with chest pain. *Ann Intern Med* 1999;130(9):709-18. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Kurita A, Chaitman BR, Bourassa MG. Significance of exercise-induced junctional S-T depression in evaluation of coronary artery disease. *Am J Cardiol* 1977;40(4):492-7. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Kurrelmeyer KM. Noninvasive evaluation of women with coronary artery disease. *Curr Opin Cardiol* 2002;17(5):464-9. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Kurt M, Shaikh KA, Peterson L,, et al. Impact of contrast echocardiography on evaluation of ventricular function and clinical management in a large prospective cohort. *J Am Coll Cardiol* 2009;53(9):802-10. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup; No outcomes of interest.*

Kwok Y, Kim C, Grady D,, et al. Meta-analysis of exercise testing to detect coronary artery disease in women. *Am J Cardiol* 1999;83(5):660-6. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Kwon SW, Kim YJ, Shim J,, et al. Coronary artery calcium scoring does not add prognostic value to standard 64-section CT angiography protocol in low-risk patients suspected of having coronary artery disease. *Radiology* 2011;259(1):92-9. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Kwong RY, Schussheim AE, Rekhraj S,, et al. Detecting acute coronary syndrome in the emergency department with cardiac magnetic resonance imaging. *Circulation* 2003;107(4):531-7. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

La Grutta L, Runza G, Gentile G,, et al. Prognostic outcome of routine clinical noninvasive multidetector-row computed tomography coronary angiography in patients with suspected coronary artery disease: a 2-year follow-up study. *Radiol Med* 2011;116(4):521-31. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

LaBounty TM, Earls JP, Leipsic J,, et al. Effect of a standardized quality-improvement protocol on radiation dose in coronary computed tomographic angiography. *Am J Cardiol* 2010;106(11):1663-7. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

LaBounty TM, Kim RJ, Lin FY,, et al. Diagnostic accuracy of coronary computed tomography angiography as interpreted on a mobile handheld phone device. *JACC Cardiovasc Imaging* 2010;3(5):482-90. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

LaBounty TM, Leipsic J, Srichai MB,, et al. What is the optimal number of readers needed to achieve high diagnostic accuracy in coronary computed tomographic angiography? A comparison of alternate reader

combinations. *J Cardiovasc Comput Tomogr* 2010;4(6):384-90. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Ladenheim ML, Pollock BH, Rozanski A, et al. Extent and severity of myocardial hypoperfusion as predictors of prognosis in patients with suspected coronary artery disease. *J Am Coll Cardiol* 1986;7(3):464-71. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

LaManna MM, Mohama R, Slavich IL, 3rd., et al. Intravenous adenosine (adenoscan) versus exercise in the noninvasive assessment of coronary artery disease by SPECT. *Clin Nucl Med* 1990;15(11):804-5. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Lamont DH, Budoff MJ, Shavelle DM, et al. Coronary calcium scanning adds incremental value to patients with positive stress tests. *Am Heart J* 2002;143(5):861-7. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Lancellotti P, Benoit T, Rigo P, et al. Dobutamine stress echocardiography versus quantitative technetium-99m sestamibi SPECT for detecting residual stenosis and multivessel disease after myocardial infarction. *Heart* 2001;86(5):510-5. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Lancellotti P, Lebois F, Simon M, et al. Prognostic importance of quantitative exercise Doppler echocardiography in asymptomatic valvular aortic stenosis. *Circulation* 2005;112(9 Suppl):I377-82. *Full-text exclusion reason(s): No data for NITs of interest (ECG, ECHO, SPECT, PET, CMR, CTA).*

Langer C, Wiemer M, Peterschroder A, et al. Stratification for noninvasive coronary angiography: patient preselection considering atypical angina pectoris, conventional cardiovascular risk assessment, and calcium scoring. *Eur J Cardiovasc Prev Rehabil* 2009;16(2):201-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Langerak SE, Vliegen HW, Jukema JW, et al. Value of magnetic resonance imaging for the noninvasive detection of stenosis in coronary artery bypass grafts and recipient coronary arteries. *Circulation* 2003;107(11):1502-8. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Lanza GA, Buffon A, Sestito A, et al. Relation between stress-induced myocardial perfusion defects on cardiovascular magnetic resonance and coronary microvascular dysfunction in patients with cardiac syndrome X. *J Am Coll Cardiol* 2008;51(4):466-72. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup; No outcomes of interest.*

Lanzarini L, Fetiveau R, Poli A, et al. Results of dipyridamole plus atropine echo stress test for the diagnosis of coronary artery disease. *Int J Card Imaging* 1995;11(4):233-40. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Laraudogoitia Zaldumbide E, Perez-David E, Larena JA, et al. The value of cardiac magnetic resonance in patients with acute coronary syndrome and normal coronary arteries. *Rev Esp Cardiol* 2009;62(9):976-83. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup; No outcomes of interest.*

Larsen J, Brekke M, Sandvik L, et al. Silent coronary atheromatosis in type 1 diabetic patients and its relation to long-term glycemic control. *Diabetes* 2002;51(8):2637-41. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Laspas F, Tsantioti D, Roussakis A, et al. Correlation of radiation dose and heart rate in dual-source computed tomography coronary angiography. *Acta Radiol* 2011;52(3):273-7. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Latcham AP, Orsinelli DA, Pearson AC. Recognition of the segmental tendency of false-positive dobutamine stress echocardiograms and its effects on test sensitivity and specificity. *Am Heart J* 1995;129(5):1047-50. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Lattanzi F, Picano E, Frugoli A, et al. Oral vs intravenous dipyridamole echocardiography for detecting coronary artery disease. *Chest* 1992;102(4):1189-92. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Lau GT, Ridley LJ, Schieb MC, et al. Coronary artery stenoses: detection with calcium scoring, CT angiography, and both methods combined. *Radiology* 2005;235(2):415-22. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Laufer E, Wahi S, Lim YL. Cost-effectiveness and accuracy of exercise stress echocardiography in the non-invasive diagnosis of coronary heart disease. *Aust N Z J Med* 2000;30(6):660-7. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Lavoie KL, Fleet RP, Lesperance F, et al. Are exercise stress tests appropriate for assessing myocardial ischemia in patients with major depressive disorder? *Am Heart J* 2004;148(4):621-7. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Law WY, Yang CC, Chen LK, et al. Retrospective gating vs. prospective triggering for noninvasive coronary angiography: Assessment of image quality and radiation dose using a 256-slice CT scanner with 270 ms gantry rotation. *Acad Radiol* 2011;18(1):31-9. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Lazoura O, Vlychou M, Vassiou K, et al. 128-detector-row computed tomography coronary angiography evaluating coronary artery disease: who avoids cardiac catheterization? *Angiology* 2010;61(2):174-8. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

le Polain de Waroux JB, Pouleur AC, Goffinet C, et al. Combined coronary and late-enhanced multidetector-computed tomography for delineation of the etiology of left ventricular dysfunction: comparison with coronary angiography and contrast-enhanced cardiac magnetic resonance imaging. *Eur Heart J* 2008;29(20):2544-51. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup; No outcomes of interest.*

Leao Lima Rde S, De Lorenzo A, Issa A. Reduced adverse effects with an accelerated dobutamine stress protocol compared with the conventional protocol: a prospective, randomized myocardial perfusion scintigraphy study. *Int J Cardiovasc Imaging* 2008;24(1):55-9. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Leber AW, Becker A, Knez A, et al. Accuracy of 64-slice computed tomography to classify and quantify plaque volumes in the proximal coronary system: a comparative study using intravascular ultrasound. *J Am Coll Cardiol* 2006;47(3):672-7. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup; No outcomes of interest.*

Leber AW, Johnson T, Becker A, et al. Diagnostic accuracy of dual-source multi-slice CT-coronary angiography in patients with an intermediate pretest likelihood for coronary artery disease. *Eur Heart J* 2007;28(19):2354-60. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Leber AW, Knez A, Becker A, et al. Accuracy of multidetector spiral computed tomography in identifying and differentiating the composition of coronary atherosclerotic plaques: a comparative study with intracoronary ultrasound. *J Am Coll Cardiol* 2004;43(7):1241-7. *Full-text exclusion reason(s): Data for women not reported as a subgroup; No outcomes of interest.*

Leber AW, Knez A, Becker C, et al. Non-invasive intravenous coronary angiography using electron beam tomography and multislice computed tomography. *Heart* 2003;89(6):633-9. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Leber AW, Knez A, Mukherjee R,, et al. Usefulness of calcium scoring using electron beam computed tomography and noninvasive coronary angiography in patients with suspected coronary artery disease. *Am J Cardiol* 2001;88(3):219-23. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Leber AW, Knez A, von Ziegler F,, et al. Quantification of obstructive and nonobstructive coronary lesions by 64-slice computed tomography: a comparative study with quantitative coronary angiography and intravascular ultrasound. *J Am Coll Cardiol* 2005;46(1):147-54. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Lee DH, Youn HJ, Choi YS,, et al. Coronary flow reserve is a comprehensive indicator of cardiovascular risk factors in subjects with chest pain and normal coronary angiogram. *Circ J* 2010;74(7):1405-14. *Full-text exclusion reason(s): Data for women not reported as a subgroup; No outcomes of interest.*

Lee DS, Jang MJ, Cheon GJ,, et al. Comparison of the cost-effectiveness of stress myocardial SPECT and stress echocardiography in suspected coronary artery disease considering the prognostic value of false-negative results. *J Nucl Cardiol* 2002;9(5):515-22. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Lee DS, Paeng JC, Lee MC. Implication of prognostically significant negative results on prone SPECT. *J Nucl Med* 2003;44(10):1641-3. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Lee HJ, Kim JS, Kim YJ,, et al. Diagnostic accuracy of 64-slice multidetector computed tomography for selecting coronary artery bypass graft surgery candidates. *J Thorac Cardiovasc Surg* 2010. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup; No outcomes of interest.*

Lee JH, Crump R, Ellestad MH. Significance of precordial T-wave increase during treadmill stress testing. *Am J Cardiol* 1995;76(17):1297-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Lee KH, Jang HJ, Lee SC,, et al. Myocardial thallium defects in apical hypertrophic cardiomyopathy are associated with a benign prognosis. Thallium defects in apical hypertrophy. *Int J Cardiovasc Imaging* 2003;19(5):381-8. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup; No outcomes of interest.*

Lee TC, O'Malley PG, Feuerstein I,, et al. The prevalence and severity of coronary artery calcification on coronary artery computed tomography in black and white subjects. *J Am Coll Cardiol* 2003;41(1):39-44. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup; No outcomes of interest.*

Legare JF, Haddad H, Barnes D,, et al. Myocardial scintigraphy correlates poorly with coronary angiography in the screening of transplant arteriosclerosis. *Can J Cardiol* 2001;17(8):866-72. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup; No outcomes of interest.*

Lehman SJ, Abbara S, Cury RC,, et al. Significance of cardiac computed tomography incidental findings in acute chest pain. *Am J Med* 2009;122(6):543-9. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup; No outcomes of interest.*

Leipsic J, Labounty TM, Heilbron B,, et al. Estimated radiation dose reduction using adaptive statistical iterative reconstruction in coronary CT angiography: the ERASIR study. *AJR Am J Roentgenol* 2010;195(3):655-60. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup; No outcomes of interest.*

Leipsic J, LaBounty TM, Mancini GB,, et al. A prospective randomized controlled trial to assess the diagnostic performance of reduced tube voltage for coronary CT angiography. *AJR Am J Roentgenol* 2011;196(4):801-6. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Leischik R, Dworrak B, Littwitz H,, et al. Prognostic significance of exercise stress echocardiography in 3329 outpatients (5-year longitudinal study). *Int J Cardiol* 2007;119(3):297-305. *Full-text exclusion reason(s): No women*

*with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Leite WA, Gil MA, Lima VC, et al. Exercise testing early after myocardial infarction: comparison with echocardiography, electrocardiographic monitoring and coronary arteriography. *Arq Bras Cardiol* 2008;90(3):176-81. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Lele RD, Luthra K, Sawant Y. Assessment of diastolic heart function—experience with 16-gated myocardial perfusion SPECT. *J Assoc Physicians India* 2008;56:763-8. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Lerakis S, Janik M, McLean DS, et al. Adenosine stress magnetic resonance imaging in women with low risk chest pain: the Emory University experience. *Am J Med Sci* 2010;339(3):216-20. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization.*

Leschka S, Alkadhi H, Plass A, et al. Accuracy of MSCT coronary angiography with 64-slice technology: first experience. *Eur Heart J* 2005;26(15):1482-7. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Leschka S, Scheffel H, Desbiolles L, et al. Combining dual-source computed tomography coronary angiography and calcium scoring: added value for the assessment of coronary artery disease. *Heart* 2008;94(9):1154-61. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Leschka S, Scheffel H, Husmann L, et al. Effect of decrease in heart rate variability on the diagnostic accuracy of 64-MDCT coronary angiography. *AJR Am J Roentgenol* 2008;190(6):1583-90. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Leschka S, Stolzmann P, Desbiolles L, et al. Diagnostic accuracy of high-pitch dual-source CT for the assessment of coronary stenoses: First experience. *Euro Radiol* 2009;19(12):2896-2903. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Lesser JR, Flygenring B, Knickelbine T, et al. Clinical utility of coronary CT angiography: coronary stenosis detection and prognosis in ambulatory patients. *Catheter Cardiovasc Interv* 2007;69(1):64-72. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Lessick J, Ghersin E, Dragu R, et al. Diagnostic accuracy of myocardial hypoenhancement on multidetector computed tomography in identifying myocardial infarction in patients admitted with acute chest pain syndrome. *J Comput Assist Tomogr* 2007;31(5):780-8. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Leta R, Carreras F, Alomar X, et al. [Non-invasive coronary angiography with 16 multidetector-row spiral computed tomography: a comparative study with invasive coronary angiography]. *Rev Esp Cardiol* 2004;57(3):217-24. *Full-text exclusion reason(s): Non-English.*

Lethen H, H PT, Kersting S, et al. Validation of noninvasive assessment of coronary flow velocity reserve in the right coronary artery. A comparison of transthoracic echocardiographic results with intracoronary Doppler flow wire measurements. *Eur Heart J* 2003;24(17):1567-75. *Full-text exclusion reason(s): No data for NITs of interest (ECG, ECHO, SPECT, PET, CMR, CTA).*

Leung DY, Lo ST, Liew CT, et al. Use of functional tests before angiography in patients with normal coronary arteries. *Int J Cardiol* 2005;104(3):326-31. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup; No outcomes of interest.*

Levine MG, McGill CC, Ahlberg AW, et al. Functional assessment with electrocardiographic gated single-photon emission computed tomography improves the ability of technetium-99m sestamibi myocardial perfusion imaging to predict myocardial viability in patients undergoing revascularization. *Am J Cardiol* 1999;83(1):1-5. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Lewis JF, McGorray SP, Pepine CJ. Assessment of women with suspected myocardial ischemia: review of findings of the Women's Ischemia Syndrome Evaluation (WISE) Study. *Curr Womens Health Rep* 2002;2(2):110-4. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Lewis WR, Ganim R, Sabapathy R. Utility of stress echocardiography in identifying significant coronary artery disease in patients with left bundle-branch block. *Crit Pathw Cardiol* 2007;6(3):127-30. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Li J, Li X, Wei M,, et al. Diagnostic accuracy of dual-source computed tomography in the detection of coronary chronic total occlusion: Comparison with invasive angiography. *African Journal of Biotechnology* 2011;10(19):3854-3858. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Liao L, Smith WTt, Tuttle RH,, et al. Prediction of death and nonfatal myocardial infarction in high-risk patients: a comparison between the Duke treadmill score, peak exercise radionuclide angiography, and SPECT perfusion imaging. *J Nucl Med* 2005;46(1):5-11. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Lim MC, Wong TW, Yaneza LO,, et al. Non-invasive detection of significant coronary artery disease with multi-section computed tomography angiography in patients with suspected coronary artery disease. *Clin Radiol* 2006;61(2):174-80. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Lim SJ, Choo KS, Park YH,, et al. Assessment of left ventricular function and volume in patients undergoing 128-slice coronary CT angiography with ECG-based maximum tube current modulation: a comparison with echocardiography. *Korean J Radiol* 2011;12(2):156-62. *Full-text exclusion reason(s): Data for women not reported as a subgroup; No outcomes of interest.*

Lima RS, De Lorenzo A, Pantoja MR,, et al. Incremental prognostic value of myocardial perfusion 99m-technetium-sestamibi SPECT in the elderly. *Int J Cardiol* 2004;93(2-3):137-43. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization.*

Limacher MC, Quinones MA, Poliner LR,, et al. Detection of coronary artery disease with exercise two-dimensional echocardiography. Description of a clinically applicable method and comparison with radionuclide ventriculography. *Circulation* 1983;67(6):1211-8. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Lin CJ, Hsu JC, Lai YJ,, et al. Diagnostic accuracy of dual-source CT coronary angiography in a population unselected for degree of coronary artery calcification and without heart rate modification. *Clin Radiol* 2010;65(2):109-17. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Lin F, Shaw LJ, Berman DS,, et al. Multidetector computed tomography coronary artery plaque predictors of stress-induced myocardial ischemia by SPECT. *Atherosclerosis* 2008;197(2):700-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup; No outcomes of interest.*

Lin FY, Saba S, Weinsaft JW,, et al. Relation of plaque characteristics defined by coronary computed tomographic angiography to ST-segment depression and impaired functional capacity during exercise treadmill testing in patients suspected of having coronary heart disease. *Am J Cardiol* 2009;103(1):50-8. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Lin GA, Dudley RA, Lucas FL,, et al. Frequency of stress testing to document ischemia prior to elective percutaneous coronary intervention. *JAMA* 2008;300(15):1765-73. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; No outcomes of interest.*

Lin SL, Chiou KR, Huang WC,, et al. Detection of coronary artery disease using real-time myocardial contrast echocardiography: a comparison with dual-isotope resting thallium-201/stress technetium-99m sestamibi single-photon emission computed tomography. *Heart Vessels* 2006;21(4):226-35. *Full-text exclusion reason(s): No women*

*with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Ling LH, Pellikka PA, Mahoney DW, et al. Atropine augmentation in dobutamine stress echocardiography: role and incremental value in a clinical practice setting. *J Am Coll Cardiol* 1996;28(3):551-7. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Lipiec P, Wejner-Mik P, Krzeminska-Pakula M, et al. Gated 99mTc-MIBI single-photon emission computed tomography for the evaluation of left ventricular ejection fraction: comparison with three-dimensional echocardiography. *Ann Nucl Med* 2008;22(8):723-6. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup; No outcomes of interest.*

Lipiec P, Wejner-Mik P, Krzeminska-Pakula M, et al. Detection of single-vessel coronary artery disease by dipyridamole stress echocardiography: no longer a problem? *Clin Physiol Funct Imaging* 2009;29(2):151-7. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD.*

Lipiec P, Wejner-Mik P, Krzeminska-Pakula M, et al. Accelerated stress real-time myocardial contrast echocardiography for the detection of coronary artery disease: comparison with 99mTc single photon emission computed tomography. *J Am Soc Echocardiogr* 2008;21(8):941-7. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Lipinski M, Do D, Froelicher V, et al. Comparison of exercise test scores and physician estimation in determining disease probability. *Arch Intern Med* 2001;161(18):2239-44. *Full-text exclusion reason(s): Data for women not reported as a subgroup; No outcomes of interest.*

Lipton JA, Nelwan SP, van Domburg RT, et al. Abnormal spatial QRS-T angle predicts mortality in patients undergoing dobutamine stress echocardiography for suspected coronary artery disease. *Coron Artery Dis* 2010;21(1):26-32. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Liu X, Cai Z, Cai Y, et al. Comparison of 16 slice multi-detector computed tomography and breath hold 3D magnetic resonance angiography in the detection of coronary stenosis. *Journal of Geriatric Cardiology* 2006;3(1):24-28. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Liu X, Zhao X, Huang J, et al. Comparison of 3D free-breathing coronary MR angiography and 64-MDCT angiography for detection of coronary stenosis in patients with high calcium scores. *AJR Am J Roentgenol* 2007;189(6):1326-32. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Lo KY, Leung KF, Chu CM, et al. Prognostic value of adenosine stress myocardial perfusion by cardiac magnetic resonance imaging in patients with known or suspected coronary artery disease. *QJM* 2011;104(5):425-32. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Lockie T, Ishida M, Perera D, et al. High-resolution magnetic resonance myocardial perfusion imaging at 3.0-Tesla to detect hemodynamically significant coronary stenoses as determined by fractional flow reserve. *J Am Coll Cardiol* 2011;57(1):70-5. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Loimaala A, Groundstroem K, Pasanen M, et al. Comparison of bicycle, heavy isometric, dipyridamole-atropine and dobutamine stress echocardiography for diagnosis of myocardial ischemia. *Am J Cardiol* 1999;84(12):1396-400. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Lokies J. Value of magnetocardiography for the non-invasive diagnosis of coronary artery disease. 30th Annual Congress of the International Society of Electrocardiology, Helsinki (Finland), 11-14 Jun 2003 (World Meeting Number 000 6973); 2003. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Lonnebakken MT, Bleie O, Strand E, et al. Myocardial contrast echocardiography in assessment of stable coronary artery disease at intermediate dobutamine-induced stress level. *Echocardiography* 2009;26(1):52-60. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*



Lorenzoni R, Cortigiani L, Magnani M, et al. Cost-effectiveness analysis of noninvasive strategies to evaluate patients with chest pain. *J Am Soc Echocardiogr* 2003;16(12):1287-91. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Lu B, Lu JG, Sun ML, et al. Comparison of diagnostic accuracy and radiation dose between prospective triggering and retrospective gated coronary angiography by dual-source computed tomography. *Am J Cardiol* 2011;107(9):1278-84. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Lu B, Shavelle DM, Mao S, et al. Improved accuracy of noninvasive electron beam coronary angiography. *Invest Radiol* 2004;39(2):73-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Luotolahti M, Saraste M, Hartiala J. Exercise echocardiography in the diagnosis of coronary artery disease. *Ann Med* 1996;28(1):73-7. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Ma Es, Yang Zg, Li Y, et al. Correlation of calcium measurement with low dose 64-slice CT and angiographic stenosis in patients with suspected coronary artery disease. *Int J Cardiol* 2010;140(2):249-252. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Maas AHM, Appelman YEA. Gender differences in coronary heart disease. *Netherlands Heart Journal* 2010;18(12):598-603. *Full-text exclusion reason(s): Not a clinical study report; Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry; Study does not compare one NIT to another or to diagnostic cardiac catheterization.*

Macaciel RM, Mesquita ET, Vivacqua R, et al. Safety, feasibility, and results of exercise testing for stratifying patients with chest pain in the emergency room. *Arq Bras Cardiol* 2003;81(2):174-81, 166-73. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Machaalany J, Yam Y, Ruddy TD, et al. Potential clinical and economic consequences of noncardiac incidental findings on cardiac computed tomography. *J Am Coll Cardiol* 2009;54(16):1533-41. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup; No outcomes of interest.*

Machecourt J, Denis B, Comet M, et al. [Comparison between the predictive value of thallium 201 myocardial scintigraphy during effort, clinical findings and the effort electrocardiogram. Study in 112 patients undergoing coronary angiography, without previous myocardial infarction]. *Arch Mal Coeur Vaiss* 1981;74(1):11-20. *Full-text exclusion reason(s): Non-English.*

Machecourt J, Longere P, Fagret D, et al. Prognostic value of thallium-201 single-photon emission computed tomographic myocardial perfusion imaging according to extent of myocardial defect. Study in 1,926 patients with follow-up at 33 months. *J Am Coll Cardiol* 1994;23(5):1096-106. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization.*

Maciejewski M, Zielinska M, Bolinska H, et al. Dobutamine stress echocardiography in symptomatic patients with nondiagnostic or negative treadmill exercise test. *Acta Cardiol* 2002;57(1):56-7. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Maddahi J, Abdulla A, Garcia EV, et al. Noninvasive identification of left main and triple vessel coronary artery disease: improved accuracy using quantitative analysis of regional myocardial stress distribution and washout of thallium-201. *J Am Coll Cardiol* 1986;7(1):53-60. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Maddahi J, Garcia EV, Berman DS, et al. Improved noninvasive assessment of coronary artery disease by quantitative analysis of regional stress myocardial distribution and washout of thallium-201. *Circulation* 1981;64(5):924-35. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Maddahi J, Van Train K, Prigent F, et al. Quantitative single photon emission computed thallium-201 tomography for detection and localization of coronary artery disease: optimization and prospective validation of a new technique.

J Am Coll Cardiol 1989;14(7):1689-99. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Madu EC. Transesophageal dobutamine stress echocardiography in the evaluation of myocardial ischemia in morbidly obese subjects. Chest 2000;117(3):657-61. *Full-text exclusion reason(s): No data for NITs of interest (ECG, ECHO, SPECT, PET, CMR, CTA).*

Maffei E, Palumbo A, Martini C., et al. Stress-ECG vs. CT coronary angiography for the diagnosis of coronary artery disease: a "real-world" experience. Radiol Med 2010;115(3):354-67. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Maffei E, Palumbo A, Martini C., et al. Diagnostic accuracy of 64-slice computed tomography coronary angiography in a large population of patients without revascularisation: registry data and review of multicentre trials. Radiol Med 2010;115(3):368-84. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Maffei E, Palumbo A, Martini C., et al. Diagnostic accuracy of computed tomography coronary angiography in a high risk symptomatic population. Acta bio-medica : Atenei Parmensis; 2010:47-53. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Maffei E, Palumbo A, Martini C., et al. Diagnostic accuracy of computed tomography coronary angiography in a high risk symptomatic population. Acta Biomedica de l'Ateneo Parmense 2010;81(1):47-53. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Maffei E, Seitun S, Martini C., et al. Prognostic value of CT coronary angiography: focus on obstructive vs. nonobstructive disease and on the presence of left main disease. Radiol Med 2011;116(1):15-31. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Maffei E, Seitun S, Romano M., et al. Computed tomography coronary angiography plaque burden in patients with suspected coronary artery disease. J Cardiovasc Med (Hagerstown) 2009;10(12):913-20. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization.*

Maffei S, Baroni M, Terrazzi M., et al. Preoperative assessment of coronary artery disease in aortic stenosis: a dipyridamole echocardiographic study. Ann Thorac Surg 1998;65(2):397-402. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Mahajan N, Polavaram L, Vankayala H., et al. Diagnostic accuracy of myocardial perfusion imaging and stress echocardiography for the diagnosis of left main and triple vessel coronary artery disease: a comparative meta-analysis. Heart 2010;96(12):956-66. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Mahenthiran J, Bangalore S, Yao SS., et al. Comparison of prognostic value of stress echocardiography versus stress electrocardiography in patients with suspected coronary artery disease. Am J Cardiol 2005;96(5):628-34. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Mahmari JJ, Boyce TM, Goldberg RK., et al. Quantitative exercise thallium-201 single photon emission computed tomography for the enhanced diagnosis of ischemic heart disease. J Am Coll Cardiol 1990;15(2):318-29. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Maintz D, Ozgun M, Hoffmeier A., et al. Whole-heart coronary magnetic resonance angiography: value for the detection of coronary artery stenoses in comparison to multislice computed tomography angiography. Acta Radiol 2007;48(9):967-73. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Mairesse GH, Marwick TH, Arnesi M., et al. Improved identification of coronary artery disease in patients with left bundle branch block by use of dobutamine stress echocardiography and comparison with myocardial perfusion tomography. Am J Cardiol 1995;76(5):321-5. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Malm S, Frigstad S, Torp H, et al. Quantitative adenosine real-time myocardial contrast echocardiography for detection of angiographically significant coronary artery disease. *J Am Soc Echocardiogr* 2006;19(4):365-72. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Maltagliati A, Berti M, Muratori M, et al. Exercise echocardiography versus exercise electrocardiography in the diagnosis of coronary artery disease in hypertension. *Am J Hypertens* 2000;13(7):796-801. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Mandysova E, Niederle P, Malkova A, et al. Usefulness of dipyridamole-handgrip echocardiography test for detecting coronary artery disease. *Am J Cardiol* 1991;67(9):883-5. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Manka R, Jahnke C, Kozerke S, et al. Dynamic 3-dimensional stress cardiac magnetic resonance perfusion imaging: detection of coronary artery disease and volumetry of myocardial hypoenhancement before and after coronary stenting. *J Am Coll Cardiol* 2011;57(4):437-44. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Manka R, Paetsch I, Schnackenburg B, et al. BOLD cardiovascular magnetic resonance at 3.0 tesla in myocardial ischemia. *J Cardiovasc Magn Reson* 2010;12:54. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Manka R, Vitonis V, Boesiger P, et al. Clinical feasibility of accelerated, high spatial resolution myocardial perfusion imaging. *JACC: Cardiovascular Imaging* 2010;3(7):710-717. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Manning WJ, Li W, Edelman RR. A preliminary report comparing magnetic resonance coronary angiography with conventional angiography. *N Engl J Med* 1993;328(12):828-32. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Mansencal N, Pellerin D, Lamar A, et al. Diagnostic value of contrast echocardiography in Tako-Tsubo cardiomyopathy. *Arch Cardiovasc Dis* 2010;103(8-9):447-53. *Full-text exclusion reason(s): All women in the study are known to have CAD; No outcomes of interest.*

Mant J, McManus RJ, Oakes RA, et al. Systematic review and modelling of the investigation of acute and chronic chest pain presenting in primary care. *Health Technol Assess* 2004;8(2):iii, 1-158. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Marangelli V, Iliceto S, Piccinni G, et al. Detection of coronary artery disease by digital stress echocardiography: comparison of exercise, transesophageal atrial pacing and dipyridamole echocardiography. *J Am Coll Cardiol* 1994;24(1):117-24. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Marano R, De Cobelli F, Floriani I, et al. Italian multicenter, prospective study to evaluate the negative predictive value of 16- and 64-slice MDCT imaging in patients scheduled for coronary angiography (NIMISCAD-Non Invasive Multicenter Italian Study for Coronary Artery Disease). *Eur Radiol* 2009;19(5):1114-23. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Marcovitz PA, Armstrong WF. Accuracy of dobutamine stress echocardiography in detecting coronary artery disease. *Am J Cardiol* 1992;69(16):1269-73. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Marie PY, Danchin N, Durand JF, et al. Long-term prediction of major ischemic events by exercise thallium-201 single-photon emission computed tomography. Incremental prognostic value compared with clinical, exercise testing, catheterization and radionuclide angiographic data. *J Am Coll Cardiol* 1995;26(4):879-86. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Mark DB, Berman DS, Budoff MJ, et al. ACCF/ACR/AHA/NASCI/SAIP/SCAI/SCCT 2010 expert consensus document on coronary computed tomographic angiography: a report of the American College of Cardiology

Foundation Task Force on Expert Consensus Documents. *Circulation* 2010;121(22):2509-43. *Full-text exclusion reason(s): Not a clinical study report.*

Marshall AJ, Hutchings F, James AJ, et al. Prognostic value of a nine-minute treadmill test in patients undergoing myocardial perfusion scintigraphy. *Am J Cardiol* 2010;106(10):1423-8. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization.*

Martin CM, McConahay DR. Maximal treadmill exercise electrocardiography. Correlations with coronary arteriography and cardiac hemodynamics. *Circulation* 1972;46(5):956-62. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Martinez-Selles M, Bueno H, Estevez A, et al. Positive non-invasive tests in the chest pain unit: importance of the clinical profile for estimating the probability of coronary artery disease. *Acute Card Care* 2008;10(4):205-8. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Martuscelli E, Razzini C, D'Eliseo A, et al. Limitations of four-slice multirow detector computed tomography in the detection of coronary stenosis. *Ital Heart J* 2004;5(2):127-31. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Martuscelli E, Romagnoli A, D'Eliseo A, et al. Accuracy of thin-slice computed tomography in the detection of coronary stenoses. *Eur Heart J* 2004;25(12):1043-8. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Maruyama T, Takada M, Hasuike T, et al. Radiation dose reduction and coronary assessability of prospective electrocardiogram-gated computed tomography coronary angiography: comparison with retrospective electrocardiogram-gated helical scan. *J Am Coll Cardiol* 2008;52(18):1450-5. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Maruyama T, Yoshizumi T, Tamura R, et al. Comparison of visibility and diagnostic capability of noninvasive coronary angiography by eight-slice multidetector-row computed tomography versus conventional coronary angiography. *Am J Cardiol* 2004;93(5):537-42. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Marwick T, Willemart B, D'Hondt AM, et al. Selection of the optimal nonexercise stress for the evaluation of ischemic regional myocardial dysfunction and malperfusion. Comparison of dobutamine and adenosine using echocardiography and 99mTc-MIBI single photon emission computed tomography. *Circulation* 1993;87(2):345-54. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Marwick TH, Case C, Sawada S, et al. Prediction of mortality using dobutamine echocardiography. *J Am Coll Cardiol* 2001;37(3):754-60. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Marwick TH, Case C, Sawada S, et al. Use of stress echocardiography to predict mortality in patients with diabetes and known or suspected coronary artery disease. *Diabetes Care* 2002;25(6):1042-8. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Marwick TH, Case C, Sawada S, et al. Prediction of outcomes in hypertensive patients with suspected coronary disease. *Hypertension* 2002;39(6):1113-8. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Marwick TH, Case C, Short L, et al. Prediction of mortality in patients without angina: use of an exercise score and exercise echocardiography. *Eur Heart J* 2003;24(13):1223-30. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Marwick TH, Case C, Vasey C, et al. Prediction of mortality by exercise echocardiography: a strategy for combination with the duke treadmill score. *Circulation* 2001;103(21):2566-71. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Marwick TH, D'Hondt AM, Mairesse GH, et al. Comparative ability of dobutamine and exercise stress in inducing myocardial ischaemia in active patients. *Br Heart J* 1994;72(1):31-8. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Marwick TH, Nemec JJ, Pashkow FJ, et al. Accuracy and limitations of exercise echocardiography in a routine clinical setting. *J Am Coll Cardiol* 1992;19(1):74-81. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Marwick TH, Nemec JJ, Stewart WJ, et al. Diagnosis of coronary artery disease using exercise echocardiography and positron emission tomography: comparison and analysis of discrepant results. *J Am Soc Echocardiogr* 1992;5(3):231-8. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Marwick TH, Shaw L, Case C, et al. Clinical and economic impact of exercise electrocardiography and exercise echocardiography in clinical practice. *Eur Heart J* 2003;24(12):1153-63. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Marwick TH, Torelli J, Harjai K, et al. Influence of left ventricular hypertrophy on detection of coronary artery disease using exercise echocardiography. *J Am Coll Cardiol* 1995;26(5):1180-6. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Mason JR, Palac RT, Freeman ML, et al. Thallium scintigraphy during dobutamine infusion: nonexercise-dependent screening test for coronary disease. *Am Heart J* 1984;107(3):481-5. *Full-text exclusion reason(s): Population does not include women  $\geq$  age 18; No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Masood Y, Liu YH, Depuey G, et al. Clinical validation of SPECT attenuation correction using x-ray computed tomography-derived attenuation maps: multicenter clinical trial with angiographic correlation. *J Nucl Cardiol* 2005;12(6):676-86. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Matsumoto N, Sato Y, Suzuki Y, et al. Usefulness of rapid low-dose/high-dose 1-day 99mTc-sestamibi ECG-gated myocardial perfusion single-photon emission computed tomography. *Circ J* 2006;70(12):1585-9. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Matsumura Y, Hozumi T, Arai K, et al. Non-invasive assessment of myocardial ischaemia using new real-time three-dimensional dobutamine stress echocardiography: comparison with conventional two-dimensional methods. *Eur Heart J* 2005;26(16):1625-32. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Matsumura Y, Hozumi T, Watanabe H, et al. Cut-off value of coronary flow velocity reserve by transthoracic Doppler echocardiography for diagnosis of significant left anterior descending artery stenosis in patients with coronary risk factors. *Am J Cardiol* 2003;92(12):1389-93. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Matsuo S, Nakajima K, Akhter N, et al. Clinical usefulness of novel cardiac MDCT/SPECT fusion image. *Ann Nucl Med* 2009;23(6):579-86. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Matsuo S, Nakamura Y, Matsumoto T, et al. Visual assessment of coronary artery stenosis with electrocardiographically-gated multislice computed tomography. *Int J Cardiovasc Imaging* 2004;20(1):61-6. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Matsushima Y, Takase B, Uehata A, et al. Comparative predictive and diagnostic value of flow-mediated vasodilation in the brachial artery and intima media thickness of the carotid artery for assessment of coronary artery disease severity. *Int J Cardiol* 2007;117(2):165-72. *Full-text exclusion reason(s): No data for NITs of interest (ECG, ECHO, SPECT, PET, CMR, CTA).*

Mattera JA, De Leon CM, Wackers FJ, et al. Association of patients' perception of health status and exercise electrocardiogram, myocardial perfusion imaging, and ventricular function measures. *Am Heart J* 2000;140(3):409-

18. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup; No outcomes of interest.*

Mazeika PK, Nadazdin A, Oakley CM. Dobutamine stress echocardiography for detection and assessment of coronary artery disease. *J Am Coll Cardiol* 1992;19(6):1203-11. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Mazur W, Rivera JM, Khoury AF, et al. Prognostic value of exercise echocardiography: validation of a new risk index combining echocardiographic, treadmill, and exercise electrocardiographic parameters. *J Am Soc Echocardiogr* 2003;16(4):318-25. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

McCarthy RM, Deshpande VS, Beohar N, et al. Three-dimensional breathhold magnetization-prepared TrueFISP: a pilot study for magnetic resonance imaging of the coronary artery disease. *Invest Radiol* 2007;42(10):665-70. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

McHenry PL, Phillips JF, Knoebel SB. Correlation of computer-quantitated treadmill exercise electrocardiogram with arteriographic location of coronary artery disease. *Am J Cardiol* 1972;30(7):747-52. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

McKeogh JR. The diagnostic role of stress echocardiography in women with coronary artery disease: evidence based review. *Curr Opin Cardiol* 2007;22(5):429-33. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

McNeill AJ, Fioretti PM, el-Said SM, et al. Enhanced sensitivity for detection of coronary artery disease by addition of atropine to dobutamine stress echocardiography. *Am J Cardiol* 1992;70(1):41-6. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Meijboom WB, Mollet NR, Van Mieghem CA, et al. 64-Slice CT coronary angiography in patients with non-ST elevation acute coronary syndrome. *Heart* 2007;93(11):1386-92. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Meijboom WB, van Mieghem CA, Mollet NR, et al. 64-slice computed tomography coronary angiography in patients with high, intermediate, or low pretest probability of significant coronary artery disease. *J Am Coll Cardiol* 2007;50(15):1469-75. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Meijboom WB, Van Mieghem CA, van Pelt N, et al. Comprehensive assessment of coronary artery stenoses: computed tomography coronary angiography versus conventional coronary angiography and correlation with fractional flow reserve in patients with stable angina. *J Am Coll Cardiol* 2008;52(8):636-43. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Meijer AB, O YL, Geleijns J, et al. Meta-analysis of 40- and 64-MDCT angiography for assessing coronary artery stenosis. *AJR Am J Roentgenol* 2008;191(6):1667-75. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Mendelson MA, Spies SM, Spies WG, et al. Usefulness of single-photon emission computed tomography of thallium-201 uptake after dipyridamole infusion for detection of coronary artery disease. *Am J Cardiol* 1992;69(14):1150-5. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Meng L, Cui L, Cheng Y, et al. Effect of heart rate and coronary calcification on the diagnostic accuracy of the dual-source CT coronary angiography in patients with suspected coronary artery disease. *Korean J Radiol* 2009;10(4):347-54. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Mennicke M, Giesler T, Ropers D, et al. [Influence of heart rate on image quality and detection of coronary stenoses with multislice spiral CT]. *Biomed Tech (Berl)* 2002;47 Suppl 1 Pt 2:782-5. *Full-text exclusion reason(s): Non-English.*

Menon M, Lesser JR, Hara H, et al. Multidetector CT coronary angiography for patient triage to invasive coronary angiography: Performance and cost in ambulatory patients with equivocal or suspected inaccurate noninvasive stress

tests. *Catheter Cardiovasc Interv* 2009;73(4):497-502. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Merhige ME, Breen WJ, Shelton V, et al. Impact of myocardial perfusion imaging with PET and (82)Rb on downstream invasive procedure utilization, costs, and outcomes in coronary disease management. *J Nucl Med* 2007;48(7):1069-76. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Merkle N, Wohrle J, Grebe O, et al. Assessment of myocardial perfusion for detection of coronary artery stenoses by steady-state, free-precession magnetic resonance first-pass imaging. *Heart* 2007;93(11):1381-5. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Merz NB, Johnson BD, Kelsey PSF, et al. Diagnostic, prognostic, and cost assessment of coronary artery disease in women. *Am J Manag Care* 2001;7(10):959-65. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Metz LD, Beattie M, Hom R, et al. The prognostic value of normal exercise myocardial perfusion imaging and exercise echocardiography: a meta-analysis. *J Am Coll Cardiol* 2007;49(2):227-37. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Meyer C, Strach K, Thomas D, et al. High-resolution myocardial stress perfusion at 3 T in patients with suspected coronary artery disease. *Eur Radiol* 2008;18(2):226-33. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Michaelides AP, Andrikopoulos GK, Antoniadis C, et al. Duration of treadmill exercise testing combined with QRS score predicts adverse cardiac outcome at long-term follow-up. *Coron Artery Dis* 2009;20(5):337-42. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Michaelides AP, Fourlas CA, Andrikopoulos GK, et al. QRS score versus ST-segment changes in patients undergoing TI-201 scintigraphy using dipyridamole infusion. *J Nucl Cardiol* 2005;12(2):203-7. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Michaelides AP, Fourlas CA, Giannopoulos N, et al. Significance of QRS duration changes in the evaluation of ST-segment depression presenting exclusively during the postexercise recovery period. *Ann Noninvasive Electrocardiol* 2006;11(3):241-6. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Michaelides AP, Massias S, Antoniadis C, et al. Novel methodology for the detection of exercise-induced myocardial wall motion abnormalities by surface electrocardiogram during exercise test. *J Electrocardiol* 2011;44(3):377-382. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Michaelides AP, Psomadaki ZD, Aigyptiadou MN, et al. Significance of exercise-induced ST changes in leads aVR, V5, and V1. Discrimination of patients with single- or multivessel coronary artery disease. *Clin Cardiol* 2003;26(5):226-30. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Michaelides AP, Tousoulis D, Fourlas CA, et al. Hypertensive patients with false-positive thallium-201 scintigraphic results in the infero-posterior wall are in high risk for coronary artery disease development. *Int J Cardiol* 2007;117(2):178-83. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Mieres JH, Rosman DR, Shaw LJ. The clinical role of stress myocardial perfusion imaging in women with suspected coronary artery disease. *Curr Cardiol Rep* 2004;6(1):27-31. *Full-text exclusion reason(s): Not a clinical study report.*

Mieres JH, Shaw LJ, Arai A, et al. Role of noninvasive testing in the clinical evaluation of women with suspected coronary artery disease: Consensus statement from the Cardiac Imaging Committee, Council on Clinical Cardiology, and the Cardiovascular Imaging and Intervention Committee, Council on Cardiovascular Radiology and

Intervention, American Heart Association. *Circulation* 2005;111(5):682-96. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Milavetz JJ, Miller TD, Hodge DO, et al. Accuracy of single-photon emission computed tomography myocardial perfusion imaging in patients with stents in native coronary arteries. *Am J Cardiol* 1998;82(7):857-61. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Miller CD, Hwang W, Hoekstra JW, et al. Stress cardiac magnetic resonance imaging with observation unit care reduces cost for patients with emergent chest pain: a randomized trial. *Ann Emerg Med* 2010;56(3):209-219 e2. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Miller DD, Younis LT, Chaitman BR, et al. Diagnostic accuracy of dipyridamole technetium 99m-labeled sestamibi myocardial tomography for detection of coronary artery disease. *J Nucl Cardiol* 1997;4(1 Pt 1):18-24. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Miller JM, Rochitte CE, Dewey M, et al. Diagnostic performance of coronary angiography by 64-row CT. *N Engl J Med* 2008;359(22):2324-36. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Miller TD, Chaliki HP, Christian TF, et al. Usefulness of worsening clinical status or exercise performance in predicting future events in patients with coronary artery disease. *Am J Cardiol* 2001;88(11):1294-7. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Miller TD, Hodge DO, Milavetz JJ, et al. A normal stress SPECT scan is an effective gatekeeper for coronary angiography. *J Nucl Cardiol* 2007;14(2):187-93. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Miller TD, Roger VL, Hodge DO, et al. Gender differences and temporal trends in clinical characteristics, stress test results and use of invasive procedures in patients undergoing evaluation for coronary artery disease. *J Am Coll Cardiol* 2001;38(3):690-7. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization.*

Milosavljevic J, Ostojic M, Marinkovic J. Dipyridamole-dobutamine stress echocardiography for the detection of myocardial ischemia in patients with hypertension. *Herz* 2005;30(3):215-222. *Full-text exclusion reason(s): All women in the study are known to have CAD.*

Min JK, Dunning A, Lin FY, et al. Age- and sex-related differences in all-cause mortality risk based on coronary computed tomography angiography findings results from the international multicenter CONFIRM (Coronary CT Angiography Evaluation for Clinical Outcomes: An International Multicenter Registry) of 23,854 patients without known coronary artery disease. *J Am Coll Cardiol* 2011;58(8):849-60. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization.*

Min JK, Dunning A, Lin FY, et al. Age- and sex-related differences in all-cause mortality risk based on coronary computed tomography angiography findings: Results from the international multicenter CONFIRM (Coronary CT Angiography Evaluation for Clinical Outcomes: An International Multicenter Registry) of 23,854 patients without known coronary artery disease. *J Am Coll Cardiol* 2011;58(8):849-860. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization.*

Min JK, Dunning A, Lin FY, et al. Rationale and design of the CONFIRM (COronary CT Angiography Evaluation For Clinical Outcomes: An InteRnational Multicenter) Registry. *J Cardiovasc Comput Tomogr* 2011;5(2):84-92. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Min JK, Feignoux J, Treutenaere J, et al. The prognostic value of multidetector coronary CT angiography for the prediction of major adverse cardiovascular events: a multicenter observational cohort study. *Int J Cardiovasc Imaging* 2010;26(6):721-8. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Min JK, Robinson M, Shaw LJ, et al. Differences in episode-based care costs for multidetector computed tomographic coronary angiography versus myocardial perfusion imaging for the diagnosis of coronary artery



disease. *J Med Econ* 2008;11(2):327-40. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup; No outcomes of interest.*

Min JK, Shaw LJ, Berman DS, et al. Costs and clinical outcomes in individuals without known coronary artery disease undergoing coronary computed tomographic angiography from an analysis of Medicare category III transaction codes. *Am J Cardiol* 2008;102(6):672-8. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup; No outcomes of interest.*

Minardi G, Di Segni M, Manzara CC, et al. Diagnostic and prognostic value of dipyridamole and dobutamine stress echocardiography in patients with Q-wave acute myocardial infarction. *Am J Cardiol* 1997;80(7):847-51. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Minoves M, Garcia A, Magrina J, et al. Evaluation of myocardial perfusion defects by means of "bull's eye" images. *Clin Cardiol* 1993;16(1):16-22. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Mir-Akbari H, Ripsweden J, Jensen J, et al. Limitations of 64-detector-row computed tomography coronary angiography: calcium and motion but not short experience. *Acta Radiol* 2009;50(2):174-80. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Mishra JP, Acio E, Heo J, et al. Impact of stress single-photon emission computed tomography perfusion imaging on downstream resource utilization. *Am J Cardiol* 1999;83(9):1401-3, A8. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup; No outcomes of interest.*

Miszalski-Jamka T, Kuntz-Hehner S, Schmidt H, et al. Myocardial contrast echocardiography enhances long-term prognostic value of supine bicycle stress two-dimensional echocardiography. *J Am Soc Echocardiogr* 2009;22(11):1220-7. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Miyazono Y, Kisanuki A, Toyonaga K, et al. Usefulness of adenosine triphosphate-atropine stress echocardiography for detecting coronary artery stenosis. *Am J Cardiol* 1998;82(3):290-4. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Modena MG. Sex differences in noninvasive diagnosis of multivessel coronary artery disease. 46th Annual Scientific Session of the American College of Cardiology, Anaheim, CA (USA), 16-19 Mar 1997 (World Meeting Number 971 0068) 1997. *Full-text exclusion reason(s): Conference abstract or trial registry posting.*

Modena MG, Rossi R, Muia N, et al. Female gender as a bias for non-invasive diagnosis of multivessel coronary artery disease. XIXth Congress of the European Society of Cardiology, Stockholm (Sweden), 24-28 August 1997 (World Meeting Number 973 0006) 1997. *Full-text exclusion reason(s): Conference abstract or trial registry posting.*

Moir S, Haluska BA, Jenkins C, et al. Incremental benefit of myocardial contrast to combined dipyridamole-exercise stress echocardiography for the assessment of coronary artery disease. *Circulation* 2004;110(9):1108-13. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Moir S, Haluska BA, Jenkins C, et al. Myocardial blood volume and perfusion reserve responses to combined dipyridamole and exercise stress: a quantitative approach to contrast stress echocardiography. *J Am Soc Echocardiogr* 2005;18(11):1187-93. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Moir S, Shaw L, Haluska B, et al. Left ventricular opacification for the diagnosis of coronary artery disease with stress echocardiography: an angiographic study of incremental benefit and cost-effectiveness. *Am Heart J* 2007;154(3):510-8. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Mollet NR, Cademartiri F, Krestin GP, et al. Improved diagnostic accuracy with 16-row multi-slice computed tomography coronary angiography. *J Am Coll Cardiol* 2005;45(1):128-32. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Mollet NR, Cademartiri F, Nieman K,, et al. Multislice spiral computed tomography coronary angiography in patients with stable angina pectoris. *J Am Coll Cardiol* 2004;43(12):2265-70. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Mollet NR, Cademartiri F, Nieman K,, et al. Noninvasive assessment of coronary plaque burden using multislice computed tomography. *Am J Cardiol* 2005;95(10):1165-9. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Mollet NR, Cademartiri F, Van Mieghem C,, et al. Adjunctive value of CT coronary angiography in the diagnostic work-up of patients with typical angina pectoris. *Eur Heart J* 2007;28(15):1872-8. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Mollet NR, Cademartiri F, van Mieghem CA,, et al. High-resolution spiral computed tomography coronary angiography in patients referred for diagnostic conventional coronary angiography. *Circulation* 2005;112(15):2318-23. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Moon JH, Park EA, Lee W,, et al. The diagnostic accuracy, image quality and radiation dose of 64-slice dual-source CT in daily practice: A single institution's experience. *Korean J Radiol* 2011;12(3):308-318. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Moon JY, Chung N, Choi BW,, et al. The utility of multi-detector row spiral CT for detection of coronary artery stenoses. *Yonsei Med J* 2005;46(1):86-94. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Morgan-Hughes GJ, Roobottom CA, Owens PE,, et al. Highly accurate coronary angiography with submillimetre, 16 slice computed tomography. *Heart* 2005;91(3):308-13. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Morganroth J, Chen CC, David D,, et al. Exercise cross-sectional echocardiographic diagnosis of coronary artery disease. *Am J Cardiol* 1981;47(1):20-6. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Morise AP, Beto R, Gupta N,, et al. Exercise QT dispersion as an independent predictor of the presence of ischemia on myocardial perfusion imaging. *Ann Noninvasive Electrocardiol* 2000;5(3):240-247. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup; No outcomes of interest.*

Motoyama S, Anno H, Sarai M,, et al. Noninvasive coronary angiography with a prototype 256-row area detector computed tomography system: comparison with conventional invasive coronary angiography. *J Am Coll Cardiol* 2008;51(7):773-5. *Full-text exclusion reason(s): Not a clinical study report.*

Mowatt G, Brazzelli M, Gemmell H,, et al. Systematic review of the prognostic effectiveness of SPECT myocardial perfusion scintigraphy in patients with suspected or known coronary artery disease and following myocardial infarction. *Nucl Med Commun* 2005;26(3):217-29. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Mowatt G, Cook JA, Hillis GS,, et al. 64-Slice computed tomography angiography in the diagnosis and assessment of coronary artery disease: systematic review and meta-analysis. *Heart* 2008;94(11):1386-93. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Mowatt G, Cummins E, Waugh N,, et al. Systematic review of the clinical effectiveness and cost-effectiveness of 64-slice or higher computed tomography angiography as an alternative to invasive coronary angiography in the investigation of coronary artery disease. *Health Technol Assess* 2008;12(17):iii-iv, ix-143. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Muhlenbruch G, Seyfarth T, Soo CS,, et al. Diagnostic value of 64-slice multi-detector row cardiac CTA in symptomatic patients. *Eur Radiol* 2007;17(3):603-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Muller H, Lerch R. Tilt table exercise echocardiography assessment in the diagnosis of coronary artery disease. *Arch Cardiovasc Dis* 2008;101(3):170-4. *Full-text exclusion reason(s): No women with symptomatic chest pain, or*

*results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Muller MF, Fleisch M, Kroeker R, et al. Proximal coronary artery stenosis: three-dimensional MRI with fat saturation and navigator echo. *J Magn Reson Imaging* 1997;7(4):644-51. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Müller-Suur R, Eriksson SV, Strandberg LE, et al. Comparison of adenosine and exercise stress test for quantitative perfusion imaging in patients on beta-blocker therapy. *Cardiology* 2001(2):112-8. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Mulvagh SL, DeMaria AN, Feinstein SB, et al. Contrast echocardiography: current and future applications. *J Am Soc Echocardiogr* 2000;13(4):331-42. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Mulvagh SL, Rakowski H, Vannan MA, et al. American Society of Echocardiography Consensus Statement on the Clinical Applications of Ultrasonic Contrast Agents in Echocardiography. *J Am Soc Echocardiogr* 2008;21(11):1179-201; quiz 1281. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Mumma BE, Baumann BM, Diercks DB, et al. Sex bias in cardiovascular testing: the contribution of patient preference. *Ann Emerg Med* 2011;57(6):551-560 e4. *Full-text exclusion reason(s): No outcomes of interest.*

Murata E, Hozumi T, Matsumura Y, et al. Coronary flow velocity reserve measurement in three major coronary arteries using transthoracic Doppler echocardiography. *Echocardiography* 2006;23(4):279-86. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Muro T, Hozumi T, Watanabe H, et al. Assessment of myocardial perfusion abnormalities by intravenous myocardial contrast echocardiography with harmonic power Doppler imaging: comparison with positron emission tomography. *Heart* 2003;89(2):145-9. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Murphy JC, Scott PJ, Shannon HJ, et al. ST elevation on the exercise ECG in patients presenting with chest pain and no prior history of myocardial infarction. *Heart* 2009;95(21):1792-7. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Murray CJ, Lopez AD. Evidence-based health policy—lessons from the Global Burden of Disease Study. *Science* 1996;274(5288):740-3. *Full-text exclusion reason(s): No data for NITs of interest (ECG, ECHO, SPECT, PET, CMR, CTA).*

Muscholl MW, Oswald M, Mayer C, et al. Prognostic value of 2D echocardiography in patients presenting with acute chest pain and non-diagnostic ECG for ST-elevation myocardial infarction. *Int J Cardiol* 2002;84(2-3):217-25. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Musto C, Simon P, Nicol E, et al. 64-multislice computed tomography in consecutive patients with suspected or proven coronary artery disease: initial single center experience. *Int J Cardiol* 2007;114(1):90-7. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Nagata M, Kato S, Kitagawa K, et al. Diagnostic accuracy of 1.5-T unenhanced whole-heart coronary MR angiography performed with 32-channel cardiac coils: initial single-center experience. *Radiology* 2011;259(2):384-92. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup; No outcomes of interest.*

Nagel E, Klein C, Paetsch I, et al. Magnetic resonance perfusion measurements for the noninvasive detection of coronary artery disease. *Circulation* 2003;108(4):432-7. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Nagel E, Lehmkuhl HB, Bocksch W, et al. Noninvasive diagnosis of ischemia-induced wall motion abnormalities with the use of high-dose dobutamine stress MRI: comparison with dobutamine stress echocardiography. *Circulation* 1999;99(6):763-70. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Nagurney JT, Bamberg F, Nichols JH, et al. The Disposition Decision on Emergency Department Patients with Chest Pain is Affected by the Results of Multi-Detector Computed Axial Tomography Scan of the Coronary Arteries. *J Emerg Med* 2010;39(1):57-64. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup; No outcomes of interest.*

Nakamura M, Takeda K, Ichihara T, et al. Feasibility of simultaneous stress 99mTc-sestamibi/rest 201Tl dual-isotope myocardial perfusion SPECT in the detection of coronary artery disease. *J Nucl Med* 1999;40(6):895-903. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Nakazato R, Tamarappoo BK, Kang X, et al. Quantitative upright-supine high-speed SPECT myocardial perfusion imaging for detection of coronary artery disease: correlation with invasive coronary angiography. *J Nucl Med* 2010;51(11):1724-31. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Nallamothu BK, Saint S, Bielak LF, et al. Electron-beam computed tomography in the diagnosis of coronary artery disease: a meta-analysis. *Arch Intern Med* 2001;161(6):833-8. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Nallamothu N, Pancholy SB, Lee KR, et al. Impact on exercise single-photon emission computed tomographic thallium imaging on patient management and outcome. *J Nucl Cardiol* 1995;2(4):334-8. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Nanasato M, Hirayama H, Ando A, et al. Incremental predictive value of myocardial scintigraphy with 123I-BMIPP in patients with acute myocardial infarction treated with primary percutaneous coronary intervention. *European journal of nuclear medicine and molecular imaging* 2004(11):1512-21. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Nance Jr JW, Bamberg F, Schoepf UJ, et al. Coronary atherosclerosis in African American and white patients with acute chest pain: Characterization with coronary CT angiography. *Radiology* 2011;260(2):373-380. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Nance JW, Jr., Bastarrika G, Kang DK, et al. High-temporal resolution dual-energy computed tomography of the heart using a novel hybrid image reconstruction algorithm: initial experience. *J Comput Assist Tomogr* 2011;35(1):119-25. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Nandalur KR, Dwamena BA, Choudhri AF, et al. Diagnostic performance of stress cardiac magnetic resonance imaging in the detection of coronary artery disease: a meta-analysis. *J Am Coll Cardiol* 2007;50(14):1343-53. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Napel S, Rubin GD, Jeffrey RB, Jr. STS-MIP: a new reconstruction technique for CT of the chest. *J Comput Assist Tomogr* 1993;17(5):832-8. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup; No outcomes of interest.*

Naser N, Buksa M, Sokolovic S, et al. The role of dobutamine stress echocardiography in detecting coronary artery disease compared with coronary angiography. *Med Arh* 2011;65(3):140-4. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Nasis A, Leung MC, Antonis PR, et al. Diagnostic accuracy of noninvasive coronary angiography with 320-detector row computed tomography. *Am J Cardiol* 2010;106(10):1429-35. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Nemes A, Balazs E, Pinter S, et al. Long-term prognostic significance of coronary flow velocity reserve in patients with significant coronary artery disease not involving the left anterior descending coronary artery (results from the SZEGED study). *Echocardiography* 2010;27(3):306-10. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Newman RJ, Darrow M, Cummings DM,, et al. Predictive value of exercise stress testing in a family medicine population. *J Am Board Fam Med* 2008;21(6):531-8. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Nguyen T, Heo J, Ogilby JD,, et al. Single photon emission computed tomography with thallium-201 during adenosine-induced coronary hyperemia: correlation with coronary arteriography, exercise thallium imaging and two-dimensional echocardiography. *J Am Coll Cardiol* 1990;16(6):1375-83. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Nicol ED, Stirrup J, Reyes E,, et al. Comparison of 64-slice cardiac computed tomography with myocardial perfusion scintigraphy for assessment of global and regional myocardial function and infarction in patients with low to intermediate likelihood of coronary artery disease. *J Nucl Cardiol* 2008;15(4):497-502. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Nicol ED, Stirrup J, Reyes E,, et al. Sixty-four-slice computed tomography coronary angiography compared with myocardial perfusion scintigraphy for the diagnosis of functionally significant coronary stenoses in patients with a low to intermediate likelihood of coronary artery disease. *J Nucl Cardiol* 2008;15(3):311-8. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Nieman K, Cademartiri F, Lemos PA,, et al. Reliable noninvasive coronary angiography with fast submillimeter multislice spiral computed tomography. *Circulation* 2002;106(16):2051-4. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Nieman K, Galema T, Weustink A,, et al. Computed tomography versus exercise electrocardiography in patients with stable chest complaints: real-world experiences from a fast-track chest pain clinic. *Heart* 2009;95(20):1669-75. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Nieman K, Galema TW, Neeffjes LA,, et al. Comparison of the value of coronary calcium detection to computed tomographic angiography and exercise testing in patients with chest pain. *Am J Cardiol* 2009;104(11):1499-504. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Nieman K, Rensing BJ, van Geuns RJ,, et al. Usefulness of multislice computed tomography for detecting obstructive coronary artery disease. *Am J Cardiol* 2002;89(8):913-8. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Nieman K, Rensing BJ, van Geuns RJ,, et al. Non-invasive coronary angiography with multislice spiral computed tomography: impact of heart rate. *Heart* 2002;88(5):470-4. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Nikolaou K, Knez A, Rist C,, et al. Accuracy of 64-MDCT in the diagnosis of ischemic heart disease. *AJR Am J Roentgenol* 2006;187(1):111-7. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Nikolaou K, Rist C, Wintersperger BJ,, et al. Clinical value of MDCT in the diagnosis of coronary artery disease in patients with a low pretest likelihood of significant disease. *AJR Am J Roentgenol* 2006;186(6):1659-68. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Nishimura S, Mahmarian JJ, Boyce TM,, et al. Quantitative thallium-201 single-photon emission computed tomography during maximal pharmacologic coronary vasodilation with adenosine for assessing coronary artery disease. *J Am Coll Cardiol* 1991;18(3):736-45. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Nixdorff U, Kufner C, Achenbach S,, et al. Head-to-head comparison of dobutamine stress echocardiography and cardiac computed tomography for the detection of significant coronary artery disease. *Cardiology* 2008;110(2):81-6. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Nkoulou R, Pazhenkottil AP, Kuest SM,, et al. Semiconductor detectors allow low-dose - Low-dose 1-day SPECT myocardial perfusion imaging. *J Nucl Med* 2011;52(8):1204-1209. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Nucifora G, Badano LP, Sarraf-Zadegan N, et al. Comparison of early dobutamine stress echocardiography and exercise electrocardiographic testing for management of patients presenting to the emergency department with chest pain. *Am J Cardiol* 2007;100(7):1068-73. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Nucifora G, Schuijf JD, van Werkhoven JM, et al. Relation between Framingham risk categories and the presence of functionally relevant coronary lesions as determined on multislice computed tomography and stress testing. *Am J Cardiol* 2009;104(6):758-63. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Nygaard TW, Gibson RS, Ryan JM, et al. Prevalence of high-risk thallium-201 scintigraphic findings in left main coronary artery stenosis: comparison with patients with multiple- and single-vessel coronary artery disease. *Am J Cardiol* 1984;53(4):462-9. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Odagiri K, Uehara A, Kurata C. Vasodilator stress impairs the left ventricular function obtained with gated single-photon emission computed tomography in patients with known or suspected coronary artery disease. *Circ J* 2010;74(12):2666-73. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup; No outcomes of interest.*

Odawara H, Hamashige N, Doi Y, et al. Noninvasive detection of significant coronary artery disease in patients with vasospastic angina. Society of Nuclear Medicine 35th Annual Meeting, San Francisco, CA (USA), 14-17 Jun 1988 (World Meeting Number 882 0432). *Full-text exclusion reason(s): Conference abstract or trial registry posting.*

Ogilby JD, Kegel JG, Heo J, et al. Correlation between hemodynamic changes and tomographic sestamibi imaging during dipyridamole-induced coronary hyperemia. *J Am Coll Cardiol* 1998;31(1):75-82. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Oguzhan A, Kisacik HL, Ozdemir K, et al. Comparison of exercise stress testing with dobutamine stress echocardiography and exercise technetium-99m isonitrile single photon emission computerized tomography for diagnosis of coronary artery disease. *Jpn Heart J* 1997;38(3):333-44. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Ohara T, Hashimoto Y, Suzuki M, et al. Early diastolic flow propagation velocity detects induced diastolic dysfunction during dobutamine stress echocardiography. *Echocardiography* 2011;28(3):335-41. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup; No outcomes of interest.*

Ohkita Y, Terasawa M, Toshima H, et al. [Exercise electrocardiography—the accuracy of Master two step test correlated with coronary arteriographic findings in angina suspected patients (author's transl)]. *Rinsho Byori* 1980;28(1):19-25. *Full-text exclusion reason(s): Non-English.*

O'Keefe JC, Edwards AC, Wiseman J, et al. Comparison of exercise electrocardiography, thallium-201 myocardial imaging and exercise gated blood pool scan in patients with suspected coronary artery disease. *Aust N Z J Med* 1983;13(1):45-50. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

O'Keefe JH, Jr., Barnhart CS, Bateman TM. Comparison of stress echocardiography and stress myocardial perfusion scintigraphy for diagnosing coronary artery disease and assessing its severity. *Am J Cardiol* 1995;75(11):25D-34D. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

O'Keefe JH, Jr., Bateman TM, Silvestri R, et al. Safety and diagnostic accuracy of adenosine thallium-201 scintigraphy in patients unable to exercise and those with left bundle branch block. *Am Heart J* 1992;124(3):614-21. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Olivetti L, Mazza G, Volpi D, et al. Multislice CT in emergency room management of patients with chest pain and medium-low probability of acute coronary syndrome. *Radiol Med* 2006;111(8):1054-63. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Olmos LI, Dakik H, Gordon R,, et al. Long-term prognostic value of exercise echocardiography compared with exercise 201Tl, ECG, and clinical variables in patients evaluated for coronary artery disease. *Circulation* 1998;98(24):2679-86. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Olson MB, Kelsey SF, Matthews K,, et al. Symptoms, myocardial ischaemia and quality of life in women: results from the NHLBI-sponsored WISE Study. *Eur Heart J* 2003;24(16):1506-14. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization.*

Olszowska M, Kostkiewicz M, Tracz W,, et al. Assessment of myocardial perfusion in patients with coronary artery disease. Comparison of myocardial contrast echocardiography and 99mTc MIBI single photon emission computed tomography. *Int J Cardiol* 2003;90(1):49-55. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Oncel D, Oncel G, Tastan A,, et al. Detection of significant coronary artery stenosis with 64-section MDCT angiography. *Eur J Radiol* 2007;62(3):394-405. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Oncel D, Oncel G, Turkoglu I. Accuracy of MR coronary angiography in the evaluation of coronary artery stenosis. *Diagn Interv Radiol* 2008;14(3):153-8. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Ong K, Chin SP, Chan WL,, et al. Feasibility and accuracy of 64-row MDCT coronary imaging from a centre with early experience: a review and comparison with established centres. *Med J Malaysia* 2005;60(5):629-36. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Ong TK, Chin SP, Liew CK,, et al. Accuracy of 64-row multidetector computed tomography in detecting coronary artery disease in 134 symptomatic patients: influence of calcification. *Am Heart J* 2006;151(6):1323 e1-6. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Ontario Health Technology Assessment. Medical Advisory Secretariat, Stress echocardiography for the diagnosis of coronary artery disease: an evidence-based analysis. *Ont Health Technol Assess Ser* [Internet]. 2010 June; 10(9) 1-61. [http://www.health.gov.on.ca/english/providers/program/mas/tech/reviews/pdf/cardiac\\_stress\\_echo\\_20100528.pdf](http://www.health.gov.on.ca/english/providers/program/mas/tech/reviews/pdf/cardiac_stress_echo_20100528.pdf). 2010. *Full-text exclusion reason(s): Not a clinical study report.*

Ostojic M, Picano E, Beleslin B,, et al. Dipyridamole-dobutamine echocardiography: a novel test for the detection of milder forms of coronary artery disease. *J Am Coll Cardiol* 1994;23(5):1115-22. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Ovrehus KA, Botker HE, Jensen JM,, et al. Influence of coronary computed tomographic angiography on patient treatment and prognosis in patients with suspected stable angina pectoris. *Am J Cardiol* 2011;107(10):1473-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Ovrehus KA, Jensen JK, Mickley HF,, et al. Comparison of usefulness of exercise testing versus coronary computed tomographic angiography for evaluation of patients suspected of having coronary artery disease. *Am J Cardiol* 2010;105(6):773-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Ovrehus KA, Munkholm H, Bottcher M,, et al. Coronary computed tomographic angiography in patients suspected of coronary artery disease: impact of observer experience on diagnostic performance and interobserver reproducibility. *J Cardiovasc Comput Tomogr* 2010;4(3):186-94. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Ozdemir K, Kisacik HL, Oguzhan A,, et al. Comparison of exercise stress testing with dobutamine stress echocardiography and radionuclide ventriculography for diagnosis of coronary artery disease. *Jpn Heart J* 1999;40(6):715-27. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Ozmen N, Yiginer O, Uz O,, et al. ST elevation in the lead aVR during exercise treadmill testing may indicate left main coronary artery disease. *Kardiol Pol* 2010;68(10):1107-11. *Full-text exclusion reason(s): Data for women not reported as a subgroup; No outcomes of interest.*

Paetsch I, Jahnke C, Barkhausen J,, et al. Detection of coronary stenoses with contrast enhanced, three-dimensional free breathing coronary MR angiography using the gadolinium-based intravascular contrast agent gadocoletic acid

(B-22956). J Cardiovasc Magn Reson 2006;8(3):509-16. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Paetsch I, Jahnke C, Wahl A,, et al. Comparison of dobutamine stress magnetic resonance, adenosine stress magnetic resonance, and adenosine stress magnetic resonance perfusion. Circulation 2004;110(7):835-42. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Pagali SR, Madaj P, Gupta M,, et al. Interobserver variations of plaque severity score and segment stenosis score in coronary arteries using 64 slice multidetector computed tomography: a substudy of the ACCURACY trial. J Cardiovasc Comput Tomogr 2010;4(5):312-8. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Paijitprapaporn P, Jongjirasiri S, Tangpagasit L,, et al. Accuracy of sixteen-slice CT scanners in detected coronary artery disease. J Med Assoc Thai 2006;89(1):72-80. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Paladugu N, Shaqra H, Blum S,, et al. Positive vasodilator stress ECG with normal myocardial perfusion imaging and its correlation with coronary angiographic findings in African Americans and Hispanics. Clin Cardiol 2010;33(10):638-42. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup; No outcomes of interest.*

Palinkas A, Toth E, Amyot R,, et al. The value of ECG and echocardiography during stress testing for identifying systemic endothelial dysfunction and epicardial artery stenosis. Eur Heart J 2002;23(20):1587-95. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Palmas W, Friedman JD, Diamond GA,, et al. Incremental value of simultaneous assessment of myocardial function and perfusion with technetium-99m sestamibi for prediction of extent of coronary artery disease. J Am Coll Cardiol 1995;25(5):1024-31. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Panmethis M, Wangsuphachart S, Rerkpattanapipat P,, et al. Detection of coronary stenoses in chronic stable angina by multi-detector CT coronary angiography. J Med Assoc Thai 2007;90(8):1573-80. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Park JW, Leithauser B, Jung F. Magnetocardiography predicts coronary artery disease in bundle-branch block patients with acute chest pain. J Electrocardiol 2007;40(1 SUPPL.):S53. *Full-text exclusion reason(s): Conference abstract or trial registry posting.*

Park JW, Leithauser B, Vrsansky M,, et al. Dobutamine stress magnetocardiography for the detection of significant coronary artery stenoses - a prospective study in comparison with simultaneous 12-lead electrocardiography. Clin Hemorheol Microcirc 2008;39(1-4):21-32. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Park MY, Choi SJ, Kim JK,, et al. Use of multidetector computed tomography for evaluating coronary artery disease in patients undergoing dialysis. Nephrology (Carlton) 2011;16(3):285-9. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Park TH, Tayan N, Takeda K,, et al. Supine bicycle echocardiography improved diagnostic accuracy and physiologic assessment of coronary artery disease with the incorporation of intermediate stages of exercise. J Am Coll Cardiol 2007;50(19):1857-63. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Parodi G, Picano E, Marcassa C,, et al. High dose dipyridamole myocardial imaging: simultaneous sestamibi scintigraphy and two-dimensional echocardiography in the detection and evaluation of coronary artery disease. Italian Group of Nuclear Cardiology. Coron Artery Dis 1999;10(3):177-84. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Parthenakis FI, Karidis KS, Zuridakis G,, et al. Left ventricular diastolic filling changes during dipyridamole-induced ischaemia. An echo-Doppler study. Coron Artery Dis 1997;8(7):449-54. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Patel MR, Albert TS, Kandzari DE,, et al. Acute myocardial infarction: safety of cardiac MR imaging after percutaneous revascularization with stents. Radiology 2006;240(3):674-80. *Full-text exclusion reason(s): All women*



*in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Patel MR, Peterson ED, Dai D., et al. Low diagnostic yield of elective coronary angiography. *N Engl J Med* 2010;362(10):886-95. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Patterson RE, Horowitz SF, Eng C., et al. Can exercise electrocardiography and thallium-201 myocardial imaging exclude the diagnosis of coronary artery disease? Bayesian analysis of the clinical limits of exclusion and indications for coronary angiography. *Am J Cardiol* 1982;49(5):1127-35. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Pattillo RW, Fuchs S, Johnson J., et al. Predictors of prognosis by quantitative assessment of coronary angiography, single photon emission computed tomography thallium imaging, and treadmill exercise testing. *Am Heart J* 1996;131(3):582-90. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Paul JF, Ohanessian A, Caussin C., et al. [Visualization of coronary tree and detection of coronary artery stenosis using 16-slice, sub-millimeter computed tomography: preliminary experience]. *Arch Mal Coeur Vaiss* 2004;97(1):31-6. *Full-text exclusion reason(s): Non-English.*

Pazhenkottil AP, Ghadri JR, Nkoulou RN., et al. Improved outcome prediction by SPECT myocardial perfusion imaging after CT attenuation correction. *J Nucl Med* 2011;52(2):196-200. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Pedrazzini GB, D'Angeli I, Vassalli G., et al. Assessment of coronary stenosis, plaque burden and remodeling by multidetector computed tomography in patients referred for suspected coronary artery disease. *J Cardiovasc Med (Hagerstown)* 2011;12(2):122-30. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Peix A, Garcia EJ, Valiente J., et al. Ischemia in women with angina and normal coronary angiograms. *Coron Artery Dis* 2007;18(5):361-6. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup; No outcomes of interest.*

Penfornis A, Zimmermann C, Boumal D., et al. Use of dobutamine stress echocardiography in detecting silent myocardial ischaemia in asymptomatic diabetic patients: a comparison with thallium scintigraphy and exercise testing. *Diabet Med* 2001;18(11):900-5. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Peng NJ, Mar GY, Liu CP., et al. Does inadequate exercise lower the accuracy of myocardial perfusion scintigraphy? *Nucl Med Commun* 2001;22(6):625-9. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Pennell DJ, Mavrogeni S, Anagnostopoulos C., et al. Thallium myocardial perfusion tomography using intravenous dipyridamole combined with maximal dynamic exercise. *Nucl Med Commun* 1993;14(11):939-45. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Pennell DJ, Underwood SR, Ell PJ., et al. Dipyridamole magnetic resonance imaging: a comparison with thallium-201 emission tomography. *Br Heart J* 1990;64(6):362-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Pennell DJ, Underwood SR, Swanton RH., et al. Dobutamine thallium myocardial perfusion tomography. *J Am Coll Cardiol* 1991;18(6):1471-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Perrier A, Roy PM, Sanchez O., et al. Multidetector-row computed tomography in suspected pulmonary embolism. *N Engl J Med* 2005;352(17):1760-8. *Full-text exclusion reason(s): No data for NITs of interest (ECG, ECHO, SPECT, PET, CMR, CTA).*

Peteiro J, Bouzas-Mosquera A, Brouillon F., et al. Treadmill exercise echocardiography as a predictor of events in patients with left ventricular hypertrophy. *Am J Hypertens* 2010;23(7):794-801. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Peteiro J, Fabregas R, Montserrat L, et al. Comparison of treadmill exercise echocardiography before and after exercise in the evaluation of patients with known or suspected coronary artery disease. *J Am Soc Echocardiogr* 1999;12(12):1073-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Peteiro J, Monserrat L, Perez R, et al. Accuracy of peak treadmill exercise echocardiography to detect multivessel coronary artery disease: comparison with post-exercise echocardiography. *Eur J Echocardiogr* 2003;4(3):182-90. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Peters RM, Shanies SA, Peters JC. Predicting extent of coronary disease: Fuzzy cluster analysis vs. Duke treadmill score. *Journal of Clinical and Basic Cardiology* 2000;3(1):39-41. *Full-text exclusion reason(s): No data for NITs of interest (ECG, ECHO, SPECT, PET, CMR, CTA).*

Pfleiderer T, Marwan M, Schepis T, et al. Characterization of culprit lesions in acute coronary syndromes using coronary dual-source CT angiography. *Atherosclerosis* 2010;211(2):437-444. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Pfleger S, Scherhag A, Latsch A, et al. Safety of dobutamine echocardiography: No signs of myocardial cell damage or activation of the coagulation system. *Disease Management and Clinical Outcomes* 2001;3(1):15-19. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Philippe L, Merino B, Blaire T, et al. Tetrofosmin early time gated post-stress single-photon emission computed tomography imaging: feasibility and potential benefits. *J Nucl Cardiol* 2011;18(1):62-72. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup; No outcomes of interest*

Phillips LM, Mieres JH. Noninvasive assessment of coronary artery disease in women: What's next? *Curr Cardiol Rep* 2010;12(2):147-154. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Pianou N, Georgakopoulos A, Anagnostopoulos CD. Positron emission tomography for risk assessment of women investigated for coronary artery disease. *Cardiology* 2011;118(3):164-167. *Full-text exclusion reason(s): Not a clinical study report.*

Picano E, Alaimo A, Chubuchny V, et al. Noninvasive pacemaker stress echocardiography for diagnosis of coronary artery disease: a multicenter study. *J Am Coll Cardiol* 2002;40(7):1305-10. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Picano E, Bedetti G, Varga A, et al. The comparable diagnostic accuracies of dobutamine-stress and dipyridamole-stress echocardiographies: a meta-analysis. *Coron Artery Dis* 2000;11(2):151-9. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Picano E, Lattanzi F, Masini M, et al. Comparison of the high-dose dipyridamole-echocardiography test and exercise two-dimensional echocardiography for diagnosis of coronary artery disease. *Am J Cardiol* 1987;59(6):539-42. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Picano E, Molinaro S, Pasanisi E. The diagnostic accuracy of pharmacological stress echocardiography for the assessment of coronary artery disease: a meta-analysis. *Cardiovasc Ultrasound* 2008;6:30. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Picano E, Morales MA, Distante A, et al. Dipyridamole-echocardiography test in angina at rest: noninvasive assessment of coronary stenosis underlying spasm. *Am Heart J* 1986;111(4):688-91. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Picano E, Parodi O, Lattanzi F, et al. Assessment of anatomic and physiological severity of single-vessel coronary artery lesions by dipyridamole echocardiography. Comparison with positron emission tomography and quantitative arteriography. *Circulation* 1994;89(2):753-61. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Pilz G, Eierle S, Heer T, et al. Negative predictive value of normal adenosine-stress cardiac MRI in the assessment of coronary artery disease and correlation with semiquantitative perfusion analysis. *J Magn Reson Imaging* 2010;32(3):615-621. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Pilz G, Heer T, Graw M, et al. Influence of small caliber coronary arteries on the diagnostic accuracy of adenosine stress cardiac magnetic resonance imaging. *Clin Res Cardiol* 2011;100(3):201-8. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Pilz G, Jeske A, Klos M, et al. Prognostic value of normal adenosine-stress cardiac magnetic resonance imaging. *Am J Cardiol* 2008;101(10):1408-12. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Pingitore A, Picano E, Colosso MQ, et al. The atropine factor in pharmacologic stress echocardiography. Echo Persantine (EPIC) and Echo Dobutamine International Cooperative (EDIC) Study Groups. *J Am Coll Cardiol* 1996;27(5):1164-70. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Pizzuto F, Voci P, Bartolomucci F, et al. Usefulness of coronary flow reserve measured by echocardiography to improve the identification of significant left anterior descending coronary artery stenosis assessed by multidetector computed tomography. *Am J Cardiol* 2009;104(5):657-64. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Plana JC, Mikati IA, Dokainish H, et al. A randomized cross-over study for evaluation of the effect of image optimization with contrast on the diagnostic accuracy of dobutamine echocardiography in coronary artery disease The OPTIMIZE Trial. *JACC Cardiovasc Imaging* 2008;1(2):145-52. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Plass A, Azemaj N, Scheffel H, et al. Accuracy of dual-source computed tomography coronary angiography: evaluation with a standardised protocol for cardiac surgeons. *Eur J Cardiothorac Surg* 2009;36(6):1011-7. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Plass A, Grunenfelder J, Leschka S, et al. Coronary artery imaging with 64-slice computed tomography from cardiac surgical perspective. *Eur J Cardiothorac Surg* 2006;30(1):109-16. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Plein S, Greenwood JP, Ridgway JP, et al. Assessment of non-ST-segment elevation acute coronary syndromes with cardiac magnetic resonance imaging. *J Am Coll Cardiol* 2004;44(11):2173-81. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Plein S, Kozerke S, Suerder D, et al. High spatial resolution myocardial perfusion cardiac magnetic resonance for the detection of coronary artery disease. *Euro Heart J* 2008;29(17):2148-2155. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Plein S, Radjenovic A, Ridgway JP, et al. Coronary artery disease: myocardial perfusion MR imaging with sensitivity encoding versus conventional angiography. *Radiology* 2005;235(2):423-30. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Pontone G, Andreini D, Ballerini G, et al. Diagnostic work-up of unselected patients with suspected coronary artery disease: complementary role of multidetector computed tomography, symptoms and electrocardiogram stress test. *Coron Artery Dis* 2007;18(4):265-74. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Pontone G, Andreini D, Quaglia C, et al. Accuracy of multidetector spiral computed tomography in detecting significant coronary stenosis in patient populations with differing pre-test probabilities of disease. *Clin Radiol* 2007;62(10):978-85. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Poprawski K, Wierchowicki M, Michalski M, et al. Usefulness of transoesophageal dipyridamole Doppler echocardiography evaluation of coronary flow reserve for detection of left anterior descending coronary artery stenosis. Comparison with dobutamine stress echocardiography and electrocardiographic exercise test. *Kardiologia Polska* 2000;52(7):21-25. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Postel T, Frick M, Feuchtnr G, et al. Role of 16-multidetector computed tomography in the assessment of coronary artery stenoses: A prospective study of consecutive patients. *Experimental and Clinical Cardiology* 2007;12(3):149-

152. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Pouleur AC, le Polain de Waroux JB, Kefer J, et al. Direct comparison of whole-heart navigator-gated magnetic resonance coronary angiography and 40- and 64-slice multidetector row computed tomography to detect the coronary artery stenosis in patients scheduled for conventional coronary angiography. *Circ Cardiovasc Imaging* 2008;1(2):114-21. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Pozzoli MM, Fioretti PM, Salustri A, et al. Exercise echocardiography and technetium-99m MIBI single-photon emission computed tomography in the detection of coronary artery disease. *Am J Cardiol* 1991;67(5):350-5. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Pracon R, Kruk M, Kepka C, et al. Epicardial adipose tissue radiodensity is independently related to coronary atherosclerosis. A multidetector computed tomography study. *Circ J* 2011;75(2):391-7. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; No outcomes of interest.*

Pratt CM, Francis MJ, Divine GW, et al. Exercise testing in women with chest pain. Are there additional exercise characteristics that predict true positive test results? *Chest* 1989;95(1):139-44. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; No outcomes of interest.*

Previtali M, Lanzarini L, Fetiveau R, et al. Comparison of dobutamine stress echocardiography, dipyridamole stress echocardiography and exercise stress testing for diagnosis of coronary artery disease. *Am J Cardiol* 1993;72(12):865-70. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Probst C, Kovacs A, Schmitz C, et al. Quantification of coronary artery stenosis with 16-slice MSCT in patients before CABG surgery: comparison to standard invasive coronary angiography. *Heart Surg Forum* 2005;8(1):E42-6. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Pugliese F, Mollet NR, Hunink MG, et al. Diagnostic performance of coronary CT angiography by using different generations of multisection scanners: single-center experience. *Radiology* 2008;246(2):384-93. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Pugliese F, Mollet NR, Runza G, et al. Diagnostic accuracy of non-invasive 64-slice CT coronary angiography in patients with stable angina pectoris. *Eur Radiol* 2006;16(3):575-82. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Pundziute G, Schuijf JD, Jukema JW, et al. Head-to-head comparison of coronary plaque evaluation between multislice computed tomography and intravascular ultrasound radiofrequency data analysis. *JACC Cardiovasc Interv* 2008;1(2):176-82. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Pundziute G, Schuijf JD, van Werkhoven JM, et al. Head-to-head comparison between bicycle exercise testing and coronary calcium score and coronary stenoses on multislice computed tomography. *Coron Artery Dis* 2009;20(4):281-7. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Qian Z, Anderson H, Marvasty I, et al. Lesion- and vessel-specific coronary artery calcium scores are superior to whole-heart Agatston and volume scores in the diagnosis of obstructive coronary artery disease. *J Cardiovasc Comput Tomogr* 2010;4(6):391-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup; No outcomes of interest.*

Quinones MA, Verani MS, Haichin RM, et al. Exercise echocardiography versus 201Tl single-photon emission computed tomography in evaluation of coronary artery disease. Analysis of 292 patients. *Circulation* 1992;85(3):1026-31. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Quyuyumi AA, Raphael MJ, Wright C,, et al. Inability of the ST segment/heart rate slope to predict accurately the severity of coronary artery disease. *Br Heart J* 1984;51(4):395-8. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Raff GL, Chinnaiyan KM, Share DA,, et al. Radiation dose from cardiac computed tomography before and after implementation of radiation dose-reduction techniques. *JAMA* 2009;301(22):2340-8. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Raff GL, Gallagher MJ, O'Neill WW,, et al. Diagnostic accuracy of noninvasive coronary angiography using 64-slice spiral computed tomography. *J Am Coll Cardiol* 2005;46(3):552-7. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Raiker K, Sinusas AJ, Wackers FJ,, et al. One-year prognosis of patients with normal planar or single-photon emission computed tomographic technetium 99m-labeled sestamibi exercise imaging. *J Nucl Cardiol* 1994;1(5 Pt 1):449-56. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Ramakrishna G, Breen JF, Mulvagh SL,, et al. Relationship between coronary artery calcification detected by electron-beam computed tomography and abnormal stress echocardiography: association and prognostic implications. *J Am Coll Cardiol* 2006;48(10):2125-31. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Ramakrishna G, Milavetz JJ, Zinsmeister AR,, et al. Effect of exercise treadmill testing and stress imaging on the triage of patients with chest pain: CHEER substudy. *Mayo Clinic Proceedings* 2005(3):322-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Ramakrishna G, Miller TD, Breen JF,, et al. Relationship and prognostic value of coronary artery calcification by electron beam computed tomography to stress-induced ischemia by single photon emission computed tomography. *Am Heart J* 2007;153(5):807-14. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Rambihar S, Abramson B. Cardiovascular imaging and noninvasive diagnosis for older adults. *Geriatrics and Aging* 2007;10(1):14-22. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Ravipati G, Aronow WS, Lai H,, et al. Comparison of sensitivity, specificity, positive predictive value, and negative predictive value of stress testing versus 64-multislice coronary computed tomography angiography in predicting obstructive coronary artery disease diagnosed by coronary angiography. *Am J Cardiol* 2008;101(6):774-5. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Regenfus M, Ropers D, Achenbach S,, et al. Noninvasive detection of coronary artery stenosis using contrast-enhanced three-dimensional breath-hold magnetic resonance coronary angiography. *J Am Coll Cardiol* 2000;36(1):44-50. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Rehn T, Griffith LS, Achuff SC,, et al. Exercise thallium-201 myocardial imaging in left main coronary artery disease: sensitive but not specific. *Am J Cardiol* 1981;48(2):217-23. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Reis G, Marcovitz PA, Leichtman AB,, et al. Usefulness of dobutamine stress echocardiography in detecting coronary artery disease in end-stage renal disease. *Am J Cardiol* 1995;75(10):707-10. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Reis SE, Holubkov R, Conrad Smith AJ,, et al. Coronary microvascular dysfunction is highly prevalent in women with chest pain in the absence of coronary artery disease: results from the NHLBI WISE study. *Am Heart J* 2001;141(5):735-41. *Full-text exclusion reason(s): No outcomes of interest.*

Rensing BJ, Surrusys PW, de Feyter PJ. CT-Based coronary angiography. *J Invasive Cardiol* 2000;12(1):23-4. *Full-text exclusion reason(s): Not a clinical study report.*

Rerkpattanapipat P, Gandhi SK, Darty SN,, et al. Feasibility to detect severe coronary artery stenoses with upright treadmill exercise magnetic resonance imaging. *Am J Cardiol* 2003;92(5):603-6. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Reynolds HR, Srichai MB, Iqbal SN, et al. Mechanisms of myocardial infarction in women without angiographically obstructive coronary artery disease. *Circulation* 2011;124(13):1414-25. *Full-text exclusion reason(s): All women in the study are known to have CAD.*

Rieber J, Huber A, Erhard I, et al. Cardiac magnetic resonance perfusion imaging for the functional assessment of coronary artery disease: a comparison with coronary angiography and fractional flow reserve. *Eur Heart J* 2006;27(12):1465-71. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Rijneke RD, Ascoop CA, Talmon JL. Clinical significance of upsloping ST segments in exercise electrocardiography. *Circulation* 1980;61(4):671-8. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Ripsweiden J, Brismar TB, Holm J, et al. Impact on image quality and radiation exposure in coronary CT angiography: 100 kVp versus 120 kVp. *Acta Radiol* 2010;51(8):903-9. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup; No outcomes of interest.*

Rixe J, Rolf A, Conradi G, et al. Detection of relevant coronary artery disease using dual-source computed tomography in a high probability patient series: comparison with invasive angiography. *Circ J* 2009;73(2):316-22. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Rizzello V, Poldermans D, Schinkel AF, et al. Long term prognostic value of myocardial viability and ischaemia during dobutamine stress echocardiography in patients with ischaemic cardiomyopathy undergoing coronary revascularisation. *Heart* 2006;92(2):239-44. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Robertson WS, Feigenbaum H, Armstrong WF, et al. Exercise echocardiography: a clinically practical addition in the evaluation of coronary artery disease. *J Am Coll Cardiol* 1983;2(6):1085-91. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Rocchi G, Fallani F, Bracchetti G, et al. Non-invasive detection of coronary artery stenosis: a comparison among power-Doppler contrast echo, 99Tc-Sestamibi SPECT and echo wall-motion analysis. *Coron Artery Dis* 2003;14(3):239-45. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Rocha-Filho JA, Blankstein R, Shturman LD, et al. Incremental value of adenosine-induced stress myocardial perfusion imaging with dual-source CT at cardiac CT angiography. *Radiology* 2010;254(2):410-9. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Rodevand O, Hogalmen G, Gudim LP, et al. Limited usefulness of non-invasive coronary angiography with 16-detector multislice computer tomography at a community hospital. *Scand Cardiovasc J* 2006;40(2):76-82. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Rodriguez O, Picano E, Fedele S, et al. Noninvasive prediction of coronary artery disease progression by comparison of serial exercise electrocardiography and dipyridamole stress echocardiography. *Int J Cardiovasc Imaging* 2002;18(2):93-9. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Rodriguez O, Picano E, Fedele S, et al. Non-invasive prediction of angiographic progression of coronary artery disease by dipyridamole-stress echocardiography. *Coron Artery Dis* 2001;12(3):197-204. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Roger VL, Jacobsen SJ, Weston SA, et al. Sex differences in evaluation and outcome after stress testing. *Mayo Clin Proc* 2002;77(7):638-45. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Study does not compare one NIT to another or to diagnostic cardiac catheterization.*

Roger VL, Pellikka PA, Oh JK,, et al. Identification of multivessel coronary artery disease by exercise echocardiography. *J Am Coll Cardiol* 1994;24(1):109-14. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Roger VL, Pellikka PA, Oh JK,, et al. Stress echocardiography. Part I. Exercise echocardiography: techniques, implementation, clinical applications, and correlations. *Mayo Clin Proc* 1995;70(1):5-15. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Roger VL, Seward JB, Bailey KR,, et al. Aortic valve resistance in aortic stenosis: Doppler echocardiographic study and surgical correlation. *Am Heart J* 1997;134(5 Pt 1):924-9. *Full-text exclusion reason(s): No data for NITs of interest (ECG, ECHO, SPECT, PET, CMR, CTA).*

Romagnoli A, Martuscelli E, Sperandio M,, et al. Role of 64-slice cardiac computed tomography in the evaluation of patients with non-ST-elevation acute coronary syndrome. *Radiol Med* 2010;115(3):341-53. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Romanens M, Gradel C, Saner H,, et al. Comparison of (99m)Tc-sestamibi lung/heart ratio, transient ischaemic dilation and perfusion defect size for the identification of severe and extensive coronary artery disease. *European Journal of Nuclear Medicine* 2001;28(7):907-910. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Romeo F, Leo R, Clementi F,, et al. Multislice computed tomography in an asymptomatic high-risk population. *Am J Cardiol* 2007;99(3):325-8. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Ropers D, Baum U, Pohle K,, et al. Detection of coronary artery stenoses with thin-slice multi-detector row spiral computed tomography and multiplanar reconstruction. *Circulation* 2003;107(5):664-6. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Ropers D, Pohle FK, Kuettner A,, et al. Diagnostic accuracy of noninvasive coronary angiography in patients after bypass surgery using 64-slice spiral computed tomography with 330-ms gantry rotation. *Circulation* 2006;114(22):2334-41; quiz 2334. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Ropers D, Rixe J, Anders K,, et al. Usefulness of multidetector row spiral computed tomography with 64- x 0.6-mm collimation and 330-ms rotation for the noninvasive detection of significant coronary artery stenoses. *Am J Cardiol* 2006;97(3):343-8. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Rosas EA, Slomka PJ, Garcia-Rojas L,, et al. Functional impact of coronary stenosis observed on coronary computed tomography angiography: Comparison with (1)(3)N-ammonia PET. *Arch Med Res* 2010;41(8):642-8. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Rozanski A, Gransar H, Hayes SW,, et al. Comparison of long-term mortality risk following normal exercise vs adenosine myocardial perfusion SPECT. *J Nucl Cardiol* 2010;17(6):999-1008. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Rozanski A, Gransar H, Wong ND,, et al. Use of coronary calcium scanning for predicting inducible myocardial ischemia: Influence of patients' clinical presentation. *J Nucl Cardiol* 2007;14(5):669-79. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Rozanski A, Gransar H, Wong ND,, et al. Clinical outcomes after both coronary calcium scanning and exercise myocardial perfusion scintigraphy. *J Am Coll Cardiol* 2007;49(12):1352-61. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Rubello D, Zanco P, Candelpergher G,, et al. Usefulness of 99mTc-MIBI stress myocardial SPECT bull's-eye quantification in coronary artery disease. *Q J Nucl Med* 1995;39(2):111-5. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Rubinshtein R, Gaspar T, Halon DA,, et al. Prevalence and extent of obstructive coronary artery disease in patients with zero or low calcium score undergoing 64-slice cardiac multidetector computed tomography for evaluation of a chest pain syndrome. *Am J Cardiol* 2007;99(4):472-5. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Rubinshtein R, Halon DA, Gaspar T., et al. Impact of 64-slice cardiac computed tomographic angiography on clinical decision-making in emergency department patients with chest pain of possible myocardial ischemic origin. *Am J Cardiol* 2007;100(10):1522-6. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Rubinshtein R, Halon DA, Gaspar T., et al. Usefulness of 64-slice cardiac computed tomographic angiography for diagnosing acute coronary syndromes and predicting clinical outcome in emergency department patients with chest pain of uncertain origin. *Circulation* 2007;115(13):1762-8. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Rubinshtein R, Halon DA, Gaspar T., et al. Cardiac computed tomographic angiography for risk stratification and prediction of late cardiovascular outcome events in patients with a chest pain syndrome. *Int J Cardiol* 2009;137(2):108-15. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Rubinshtein R, Halon DA, Gaspar T., et al. Usefulness of 64-slice multidetector computed tomography in diagnostic triage of patients with chest pain and negative or nondiagnostic exercise treadmill test result. *Am J Cardiol* 2007;99(7):925-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Rubinshtein R, Miller TD, Williamson EE., et al. Detection of myocardial infarction by dual-source coronary computed tomography angiography using quantitated myocardial scintigraphy as the reference standard. *Heart* 2009;95(17):1419-22. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup; No outcomes of interest.*

Rumberger JA, Behrenbeck T, Breen JF., et al. Coronary calcification by electron beam computed tomography and obstructive coronary artery disease: a model for costs and effectiveness of diagnosis as compared with conventional cardiac testing methods. *J Am Coll Cardiol* 1999;33(2):453-62. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Rumberger JA, Sheedy PF, 3rd, Breen JF., et al. Coronary calcium, as determined by electron beam computed tomography, and coronary disease on arteriogram. Effect of patient's sex on diagnosis. *Circulation* 1995;91(5):1363-7. *Full-text exclusion reason(s): All women in the study are known to have CAD; No outcomes of interest.*

Russo V, Gostoli V, Lovato L., et al. Clinical value of multidetector CT coronary angiography as a preoperative screening test before non-coronary cardiac surgery. *Heart* 2007;93(12):1591-8. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Russo V, Zavalloni A, Bacchi Reggiani ML., et al. Incremental prognostic value of coronary CT angiography in patients with suspected coronary artery disease. *Circ Cardiovasc Imaging* 2010;3(4):351-9. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Ruzsics B, Lee H, Zwerner PL., et al. Dual-energy CT of the heart for diagnosing coronary artery stenosis and myocardial ischemia-initial experience. *Eur Radiol* 2008;18(11):2414-24. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Ruzsics B, Schwarz F, Schoepf UJ., et al. Comparison of dual-energy computed tomography of the heart with single photon emission computed tomography for assessment of coronary artery stenosis and of the myocardial blood supply. *Am J Cardiol* 2009;104(3):318-26. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Ryan T, Segar DS, Sawada SG., et al. Detection of coronary artery disease with upright bicycle exercise echocardiography. *J Am Soc Echocardiogr* 1993;6(2):186-97. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*



Ryan T, Vasey CG, Presti CF, et al. Exercise echocardiography: detection of coronary artery disease in patients with normal left ventricular wall motion at rest. *J Am Coll Cardiol* 1988;11(5):993-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Sa MI, Nicol ED, Stirrup J, et al. Implications for single phase prospective CT coronary angiography for the diagnosis of significant coronary stenoses in clinical practice. *Int J Cardiol* 2011;147(3):393-397. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup; No outcomes of interest.*

Saad MAM, Azer HY. Dual-source CT coronary angiography: Diagnostic accuracy without the use of B blockers. *Egyptian Journal of Radiology and Nuclear Medicine* 2011. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Sabharwal NK, Lahiri A. Role of myocardial perfusion imaging for risk stratification in suspected or known coronary artery disease. *Heart* 2003;89(11):1291-7. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Sabharwal NK, Stoykova B, Taneja AK, et al. A randomized trial of exercise treadmill ECG versus stress SPECT myocardial perfusion imaging as an initial diagnostic strategy in stable patients with chest pain and suspected CAD: cost analysis. *J Nucl Cardiol* 2007;14(2):174-86. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Sahin M, Karakelleoglu S, Alp N, et al. Diagnostic value of dobutamine stress echocardiography in coronary artery disease. *Thorac Cardiovasc Surg* 1994;42(5):285-9. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Sakakura K, Yasu T, Kobayashi Y, et al. Noninvasive tissue characterization of coronary arterial plaque by 16-slice computed tomography in acute coronary syndrome. *Angiology* 2006;57(2):155-60. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Sakuma H, Ichikawa Y, Chino S, et al. Detection of coronary artery stenosis with whole-heart coronary magnetic resonance angiography. *J Am Coll Cardiol* 2006;48(10):1946-50. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Sakuma H, Ichikawa Y, Suzawa N, et al. Assessment of coronary arteries with total study time of less than 30 minutes by using whole-heart coronary MR angiography. *Radiology* 2005;237(1):316-21. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Sakuma H, Suzawa N, Ichikawa Y, et al. Diagnostic accuracy of stress first-pass contrast-enhanced myocardial perfusion MRI compared with stress myocardial perfusion scintigraphy. *AJR Am J Roentgenol* 2005;185(1):95-102. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Salm LP, Schuijf JD, de Roos A, et al. Global and regional left ventricular function assessment with 16-detector row CT: comparison with echocardiography and cardiovascular magnetic resonance. *Eur J Echocardiogr* 2006;7(4):308-14. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup; No outcomes of interest.*

Salustri A, Fioretti PM, McNeill AJ, et al. Pharmacological stress echocardiography in the diagnosis of coronary artery disease and myocardial ischaemia: a comparison between dobutamine and dipyridamole. *Eur Heart J* 1992;13(10):1356-62. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Salustri A, Pozzoli MM, Hermans W, et al. Relationship between exercise echocardiography and perfusion single-photon emission computed tomography in patients with single-vessel coronary artery disease. *Am Heart J* 1992;124(1):75-83. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Samady H, Wackers FJ, Joska TM, et al. Pharmacologic stress perfusion imaging with adenosine: role of simultaneous low-level treadmill exercise. *J Nucl Cardiol* 2002;9(2):188-96. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in*

*the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup; No outcomes of interest.*

Sampson UK, Dorbala S, Limaye A, et al. Diagnostic accuracy of rubidium-82 myocardial perfusion imaging with hybrid positron emission tomography/computed tomography in the detection of coronary artery disease. *J Am Coll Cardiol* 2007;49(10):1052-8. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

San Roman JA, Sanz-Ruiz R, Ortega JR, et al. Safety and predictors of complications with a new accelerated dobutamine stress echocardiography protocol. *J Am Soc Echocardiogr* 2008;21(1):53-7. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

San Roman JA, Vilacosta I, Castillo JA, et al. Dipyridamole and dobutamine-atropine stress echocardiography in the diagnosis of coronary artery disease. Comparison with exercise stress test, analysis of agreement, and impact of antianginal treatment. *Chest* 1996;110(5):1248-54. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Sandstede JJ, Pabst T, Wacker C, et al. Breath-hold 3D MR coronary angiography with a new intravascular contrast agent (feruglose)—first clinical experiences. *Magn Reson Imaging* 2001;19(2):201-5. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Santana CA, Folks RD, Garcia EV, et al. Quantitative (82)Rb PET/CT: development and validation of myocardial perfusion database. *J Nucl Med* 2007;48(7):1122-8. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Santana CA, Garcia EV, Faber TL, et al. Diagnostic performance of fusion of myocardial perfusion imaging (MPI) and computed tomography coronary angiography. *J Nucl Cardiol* 2009;16(2):201-11. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Santana CA, Garcia EV, Vansant JP, et al. Gated stress-only 99mTc myocardial perfusion SPECT imaging accurately assesses coronary artery disease. *Nucl Med Commun* 2003;24(3):241-9. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Santoro GM, Sciagra R, Buonamici P, et al. Head-to-head comparison of exercise stress testing, pharmacologic stress echocardiography, and perfusion tomography as first-line examination for chest pain in patients without history of coronary artery disease. *J Nucl Cardiol* 1998;5(1):19-27. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Sarasso G, Airoidi L, Piccinino C, et al. Detection of coronary artery disease in hypertensive patients with chest pain: Diagnostic accuracy of noninvasive techniques. *Cardiovascular Imaging* 1999;11(3):133-136. *Full-text exclusion reason(s): Full-text unobtainable.*

Saraste M, Vesalainen RK, Ylitalo A, et al. Transthoracic Doppler echocardiography as a noninvasive tool to assess coronary artery stenoses—a comparison with quantitative coronary angiography. *J Am Soc Echocardiogr* 2005;18(6):679-85. *Full-text exclusion reason(s): No data for NITs of interest (ECG, ECHO, SPECT, PET, CMR, CTA).*

Sardanelli F, Molinari G, Zandrino F, et al. Three-dimensional, navigator-echo MR coronary angiography in detecting stenoses of the major epicardial vessels, with conventional coronary angiography as the standard of reference. *Radiology* 2000;214(3):808-14. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Sarno G, Decraemer I, Vanhoenacker PK, et al. On the inappropriateness of noninvasive multidetector computed tomography coronary angiography to trigger coronary revascularization: a comparison with invasive angiography. *JACC Cardiovasc Interv* 2009;2(6):550-7. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Sarullo FM, Di Pasquale P, Orlando G,, et al. Utility and safety of immediate exercise testing of low-risk patients admitted to the hospital with acute chest pain. *Int J Cardiol* 2000;75(2-3):239-43. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Sato A, Hiroe M, Tamura M,, et al. Quantitative measures of coronary stenosis severity by 64-Slice CT angiography and relation to physiologic significance of perfusion in nonobese patients: comparison with stress myocardial perfusion imaging. *J Nucl Med* 2008;49(4):564-72. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Sato A, Nozato T, Hikita H,, et al. Incremental value of combining 64-slice computed tomography angiography with stress nuclear myocardial perfusion imaging to improve noninvasive detection of coronary artery disease. *J Nucl Cardiol* 2010;17(1):19-26. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Sato Y, Matsumoto N, Ichikawa M,, et al. Efficacy of multislice computed tomography for the detection of acute coronary syndrome in the emergency department. *Circ J* 2005;69(9):1047-51. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Sato Y, Matsumoto N, Kato M,, et al. Noninvasive assessment of coronary artery disease by multislice spiral computed tomography using a new retrospectively ECG-gated image reconstruction technique. *Circ J* 2003;67(5):401-5. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Sawada SG, Ryan T, Conley MJ,, et al. Prognostic value of a normal exercise echocardiogram. *Am Heart J* 1990;120(1):49-55. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization.*

Sawada SG, Syyed S, Raiesdana A,, et al. Clinical assessment and rest and stress echocardiography for prediction of long-term prognosis in African Americans with known or suspected coronary artery disease. *Echocardiography* 2009;26(5):558-66. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Sawada SG, Segar DS, Ryan T,, et al. Echocardiographic detection of coronary artery disease during dobutamine infusion. *Circulation* 1991;83(5):1605-14. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Scalise F, Bertella M, Manfredi M,, et al. Stress-induced QTc-interval shortening as an ancillary marker of ischemia in patients with complete left bundle branch block. *J Cardiovasc Med (Hagerstown)* 2009;10(5):376-82. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Scandrett RM, Mukherjee SK. Frontiers in women's cardiovascular health prevention: What have we learned so far? *World Heart Journal* 2008;1(2):141-159. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Scanlon PJ, Faxon DP, Audet AM,, et al. ACC/AHA guidelines for coronary angiography. A report of the American College of Cardiology/American Heart Association Task Force on practice guidelines (Committee on Coronary Angiography). Developed in collaboration with the Society for Cardiac Angiography and Interventions. *J Am Coll Cardiol* 1999;33(6):1756-824. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Schannwell CM. Echocardiographic diastolic function: A sensitive noninvasive approach for the detection of coronary artery disease in women. 46th Annual Scientific Session of the American College of Cardiology, Anaheim, CA (USA), 16-19 Mar 1997 (World Meeting Number 971 0068) 1997. *Full-text exclusion reason(s): No outcomes of interest.*

Schartl M, Beckmann S, Bocksch W,, et al. Stress echocardiography in special groups: In women, in left bundle branch block, in hypertension and after heart transplantation. *Euro Heart J* 1997;18(SUPPL. D). *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Scheffel H, Alkadhi H, Leschka S, et al. Low-dose CT coronary angiography in the step-and-shoot mode: diagnostic performance. *Heart* 2008;94(9):1132-7. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Scheffel H, Alkadhi H, Plass A, et al. Accuracy of dual-source CT coronary angiography: First experience in a high pre-test probability population without heart rate control. *Eur Radiol* 2006;16(12):2739-47. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Scheffel H, Stolzmann P, Alkadhi H, et al. Low-dose CT and cardiac MR for the diagnosis of coronary artery disease: accuracy of single and combined approaches. *Int J Cardiovasc Imaging* 2010;26(5):579-90. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Scheike M, Nilsson S, Nylander E. Exercise testing and myocardial perfusion scintigraphy in primary care patients with chest pain of new onset. *Scand J Prim Health Care* 2007;25(2):117-22. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Schenker MP, Dorbala S, Hong EC, et al. Interrelation of coronary calcification, myocardial ischemia, and outcomes in patients with intermediate likelihood of coronary artery disease: a combined positron emission tomography/computed tomography study. *Circulation* 2008;117(13):1693-700. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Schepis T, Gaemperli O, Koepfli P, et al. Added value of coronary artery calcium score as an adjunct to gated SPECT for the evaluation of coronary artery disease in an intermediate-risk population. *J Nucl Med* 2007;48(9):1424-30. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Scherhag AW, Pflieger S, Schreckenberger AB, et al. Detection of patients with restenosis after PTCA by dipyridamole-atropine-stress-echocardiography. *Int J Card Imaging* 1997;13(2):115-23. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Schicha H, Rentrop P, Facorro L, et al. [Results of quantitative myocardial scintigraphy with thallium-201 at rest and after maximum exercise—critical analysis of predictive value and clinical application (author's transl)]. *Z Kardiol* 1980;69(1):31-42. *Full-text exclusion reason(s): Non-English.*

Schillaci O, Moroni C, Scopinaro F, et al. Technetium-99m sestamibi myocardial tomography based on dipyridamole echocardiography testing in hypertensive patients with chest pain. *Eur J Nucl Med* 1997;24(7):774-8. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Schinkel AF, Elhendy A, Biagini E, et al. Prognostic stratification using dobutamine stress 99mTc-tetrofosmin myocardial perfusion SPECT in elderly patients unable to perform exercise testing. *J Nucl Med* 2005;46(1):12-8. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Schinkel AF, Elhendy A, van Domburg RT, et al. Prognostic value of dobutamine-atropine stress (99m)Tc-tetrofosmin myocardial perfusion SPECT in patients with known or suspected coronary artery disease. *J Nucl Med* 2002;43(6):767-72. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Schinkel AF, Elhendy A, Van Domburg RT, et al. Long-term prognostic value of dobutamine stress 99mTc-sestamibi SPECT: single-center experience with 8-year follow-up. *Radiology* 2002;225(3):701-6. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Schinkel AF, Elhendy A, van Domburg RT, et al. Incremental value of exercise technetium-99m tetrofosmin myocardial perfusion single-photon emission computed tomography for the prediction of cardiac events. *Am J Cardiol* 2003;91(4):408-11. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Schlegel TT, Kulecz WB, Feiveson AH, et al. Accuracy of advanced versus strictly conventional 12-lead ECG for detection and screening of coronary artery disease, left ventricular hypertrophy and left ventricular systolic dysfunction. *BMC Cardiovascular Disorders* 2010;10(28). *Full-text exclusion reason(s): No data for NITs of interest (ECG, ECHO, SPECT, PET, CMR, CTA).*

Schlett CL, Banerji D, Siegel E,, et al. Prognostic value of CT angiography for major adverse cardiac events in patients with acute chest pain from the emergency department: 2-Year outcomes of the ROMICAT trial. *JACC: Cardiovascular Imaging* 2011;4(5):481-491. *Full-text exclusion reason(s): Data for women not reported as a subgroup; No outcomes of interest.*

Schlosser T, Mohrs OK, Magedanz A,, et al. Noninvasive coronary angiography using 64-detector-row computed tomography in patients with a low to moderate pretest probability of significant coronary artery disease. *Acta Radiol* 2007;48(3):300-7. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Schmermund A, Baumgart D, Sack S,, et al. Assessment of coronary calcification by electron-beam computed tomography in symptomatic patients with normal, abnormal or equivocal exercise stress test. *Eur Heart J* 2000;21(20):1674-82. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Schmermund A, Elsasser A, Behl M,, et al. Comparison of prognostic usefulness (three years) of computed tomographic angiography versus 64-slice computed tomographic calcium scanner in subjects without significant coronary artery disease. *Am J Cardiol* 2010;106(11):1574-9. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup*

Schmermund A, Stang A, Mohlenkamp S,, et al. Prognostic value of electron-beam computed tomography-derived coronary calcium scores compared with clinical parameters in patients evaluated for coronary artery disease. Prognostic value of EBCT in symptomatic patients. *Z Kardiol* 2004;93(9):696-705. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Schmidt A, Stefenelli T, Schuster E,, et al. Informational contribution of noninvasive screening tests for coronary artery disease in patients on chronic renal replacement therapy. *Am J Kidney Dis* 2001;37(1):56-63. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Schnapauff D, Dubel HP, Scholze J,, et al. Multislice computed tomography: angiographic emulation versus standard assessment for detection of coronary stenoses. *Eur Radiol* 2007;17(7):1858-64. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Schoenhagen P, Tuzcu EM, Stillman AE,, et al. Non-invasive assessment of plaque morphology and remodeling in mildly stenotic coronary segments: comparison of 16-slice computed tomography and intravascular ultrasound. *Coron Artery Dis* 2003;14(6):459-62. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Schonenberger E, Schnapauff D, Teige F,, et al. Patient acceptance of noninvasive and invasive coronary angiography. *PLoS One* 2007;2(2):e246. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup; No outcomes of interest.*

Schroder K, Voller H, Dingerkus H,, et al. Comparison of the diagnostic potential of four echocardiographic stress tests shortly after acute myocardial infarction: submaximal exercise, transesophageal atrial pacing, dipyridamole, and dobutamine-atropine. *Am J Cardiol* 1996;77(11):909-14. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup*

Schroder K, Wieckhorst A, Voller H. Comparison of the prognostic value of dipyridamole and dobutamine stress echocardiography in patients with known or suspected coronary artery disease. *Am J Cardiol* 1997;79(11):1516-8. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Schroeder S, Kuettner A, Beck T,, et al. Usefulness of noninvasive MSCT coronary angiography as first-line imaging technique in patients with chest pain: initial clinical experience. *Int J Cardiol* 2005;102(3):469-75. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Schuijff JD, Bax JJ. Noninvasive coronary angiography with multislice computed tomography and myocardial perfusion imaging. *Cardiol Rev* 2007;24(6):38-41. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Schuijf JD, Bax JJ, Jukema JW,, et al. Assessment of left ventricular volumes and ejection fraction with 16-slice multi-slice computed tomography; comparison with 2D-echocardiography. *Int J Cardiol* 2007;116(2):201-5. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup; No outcomes of interest.*

Schuijf JD, Bax JJ, Jukema JW,, et al. Noninvasive angiography and assessment of left ventricular function using multislice computed tomography in patients with type 2 diabetes. *Diabetes Care* 2004;27(12):2905-10. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Schuijf JD, Bax JJ, Salm LP,, et al. Noninvasive coronary imaging and assessment of left ventricular function using 16-slice computed tomography. *Am J Cardiol* 2005;95(5):571-4. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Schuijf JD, Mollet NR, Cademartiri F,, et al. Do risk factors influence the diagnostic accuracy of noninvasive coronary angiography with multislice computed tomography? *J Nucl Cardiol* 2006;13(5):635-41. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Schuijf JD, Pundziute G, Bax JJ. Prognostic value of multislice computed tomography coronary angiography. *Cardiol Rev* 2007;24(12):39-42. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Schuijf JD, Pundziute G, Jukema JW,, et al. Diagnostic accuracy of 64-slice multislice computed tomography in the noninvasive evaluation of significant coronary artery disease. *Am J Cardiol* 2006;98(2):145-8. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Schuijf JD, van Werkhoven JM, Pundziute G,, et al. Invasive versus noninvasive evaluation of coronary artery disease. *JACC Cardiovasc Imaging* 2008;1(2):190-9. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Schuijf JD, Wijns W, Jukema JW,, et al. Relationship between noninvasive coronary angiography with multi-slice computed tomography and myocardial perfusion imaging. *J Am Coll Cardiol* 2006;48(12):2508-14. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Schuijf JD, Wijns W, Jukema JW,, et al. A comparative regional analysis of coronary atherosclerosis and calcium score on multislice CT versus myocardial perfusion on SPECT. *J Nucl Med* 2006;47(11):1749-55. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Schutzenberger W, Leisch F, Herbinger W. [R amplitude changes and ST segment lowering in the exercise ECG compared with angiographic findings (author's transl)]. *Dtsch Med Wochenschr* 1980;105(37):1285-9. *Full-text exclusion reason(s): Non-English.*

Schwartz JG, Johnson RB, Aepfelbacher FC,, et al. Sensitivity, specificity and accuracy of stress SPECT myocardial perfusion imaging for detection of coronary artery disease in the distribution of first-order branch vessels, using an anatomical matching of angiographic and perfusion data. *Nucl Med Commun* 2003;24(5):543-9. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Schwartz L, Overgaard CB. The accuracy of noninvasive stress myocardial imaging for detecting coronary artery disease in clinical practice. *Hosp Pract (Minneap)* 2010;38(2):14-8. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Schwitzer J, Nanz D, Kneifel S,, et al. Assessment of myocardial perfusion in coronary artery disease by magnetic resonance: a comparison with positron emission tomography and coronary angiography. *Circulation* 2001;103(18):2230-5. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Schwitzer J, Wacker CM, van Rossum AC,, et al. MR-IMPACT: comparison of perfusion-cardiac magnetic resonance with single-photon emission computed tomography for the detection of coronary artery disease in a

multicentre, multivendor, randomized trial. *Eur Heart J* 2008;29(4):480-9. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Sciagra R, Zoccarato O, Bisi G, et al. Decreased [<sup>99m</sup>Tc]Sestamibi uptake with dobutamine versus dipyridamole stress. *Q J Nucl Med Mol Imaging* 2009;53(6):671-7. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Sciammarella MG, Fragasso G, Gerundini P, et al. <sup>99</sup>Tcm-MIBI single photon emission tomography (SPET) for detecting myocardial ischaemia and necrosis in patients with significant coronary artery disease. *Nucl Med Commun* 1992;13(12):871-8. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Seeck A, Garde A, Schuepbach M, et al. Diagnosis of ischemic heart disease with cardiogoniometry - Linear discriminant analysis versus support vector machines. *IFMBE Proceedings* 2008;22:389-392. *Full-text exclusion reason(s): Conference abstract or trial registry posting.*

Seeger A, Doesch C, Klumpp B, et al. [MR stress perfusion for the detection of flow-limiting stenoses in symptomatic patients with known coronary artery disease and history of stent implantation]. *Rofo* 2007;179(10):1068-73. *Full-text exclusion reason(s): Non-English.*

Seese B, Moshage W, Achenbach S, et al. Diagnostic value of coronary calcifications for non-invasive detection of coronary artery disease: A comparison of electron beam tomography, thoracic fluoroscopy and coronary angiography. XVIIth Congress of the European Society of Cardiology, Amsterdam (The Netherlands), 20-24 Aug 1995 (World Meeting Number 953 0126) 1995. *Full-text exclusion reason(s): Conference abstract or trial registry posting.*

Segar DS, Brown SE, Sawada SG, et al. Dobutamine stress echocardiography: correlation with coronary lesion severity as determined by quantitative angiography. *J Am Coll Cardiol* 1992;19(6):1197-202. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Selcoki Y, Yilmaz OC, Kankilic MN, et al. Diagnostic accuracy of 64-slice computed tomography in patients with suspected or proven coronary artery disease. *Turk Kardiyol Dern Ars* 2010;38(2):95-100. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Senior R, Basu S, Handler C, et al. Diagnostic accuracy of dobutamine stress echocardiography for detection of coronary heart disease in hypertensive patients. *Eur Heart J* 1996;17(2):289-95. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Senior R, Becher H, Monaghan M, et al. Contrast echocardiography: evidence-based recommendations by European Association of Echocardiography. *Eur J Echocardiogr* 2009;10(2):194-212. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Senior R, Janardhanan R, Jeetley P, et al. Myocardial contrast echocardiography for distinguishing ischemic from nonischemic first-onset acute heart failure: insights into the mechanism of acute heart failure. *Circulation* 2005;112(11):1587-93. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Senior R, Khattar R, Lahiri A. Value of dobutamine stress echocardiography for the detection of multivessel coronary artery disease. *Am J Cardiol* 1998;81(3):298-301. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Senior R, Lepper W, Pasquet A, et al. Myocardial perfusion assessment in patients with medium probability of coronary artery disease and no prior myocardial infarction: comparison of myocardial contrast echocardiography with <sup>99m</sup>Tc single-photon emission computed tomography. *Am Heart J* 2004;147(6):1100-5. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Senior R, Monaghan M, Main ML, et al. Detection of coronary artery disease with perfusion stress echocardiography using a novel ultrasound imaging agent: two Phase 3 international trials in comparison with radionuclide perfusion imaging. *Eur J Echocardiogr* 2009;10(1):26-35. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Senior R, Sridhara BS, Anagnostou E,, et al. Synergistic value of simultaneous stress dobutamine sestamibi single-photon-emission computerized tomography and echocardiography in the detection of coronary artery disease. *Am Heart J* 1994;128(4):713-8. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Sensky PR, Jivan A, Hudson NM,, et al. Coronary artery disease: combined stress MR imaging protocol-one-stop evaluation of myocardial perfusion and function. *Radiology* 2000;215(2):608-14. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Sensky PR, Samani NJ, Reek C,, et al. Magnetic resonance perfusion imaging in patients with coronary artery disease: a qualitative approach. *Int J Cardiovasc Imaging* 2002;18(5):373-83; discussion 385-6. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Seo WW, Chang HJ, Cho I,, et al. The value of brachial-ankle pulse wave velocity as a predictor of coronary artery disease in high-risk patients. *Korean Circulation Journal* 2010;40(5):224-229. *Full-text exclusion reason(s): No data for NITs of interest (ECG, ECHO, SPECT, PET, CMR, CTA).*

Shabestari AA, Abdi S, Akhlaghpour S,, et al. Diagnostic performance of 64-channel multislice computed tomography in assessment of significant coronary artery disease in symptomatic subjects. *Am J Cardiol* 2007;99(12):1656-61. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Shah DJ, Judd RM, Kim RJ. Technology insight: MRI of the myocardium. *Nat Clin Pract Cardiovasc Med* 2005;2(11):597-605; quiz 606. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Shaheen J, Luria D, Klutstein MW,, et al. Diagnostic value of 12-lead electrocardiogram during dobutamine echocardiographic studies. *Am Heart J* 1998;136(6):1061-4. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Shapiro MD, Butler J, Rieber J,, et al. Analytic approaches to establish the diagnostic accuracy of coronary computed tomography angiography as a tool for clinical decision making. *Am J Cardiol* 2007;99(8):1122-7. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Sharir T, Bacher-Stier C, Dhar S,, et al. Identification of severe and extensive coronary artery disease by postexercise regional wall motion abnormalities in Tc-99m sestamibi gated single-photon emission computed tomography. *Am J Cardiol* 2000;86(11):1171-5. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Sharir T, Kang X, Germano G,, et al. Prognostic value of poststress left ventricular volume and ejection fraction by gated myocardial perfusion SPECT in women and men: gender-related differences in normal limits and outcomes. *J Nucl Cardiol* 2006;13(4):495-506. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization.*

Sharp SM, Sawada SG, Segar DS,, et al. Dobutamine stress echocardiography: detection of coronary artery disease in patients with dilated cardiomyopathy. *J Am Coll Cardiol* 1994;24(4):934-9. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Sharples L, Hughes V, Crean A,, et al. Cost-effectiveness of functional cardiac testing in the diagnosis and management of coronary artery disease: a randomised controlled trial. The CECaT trial. *Health Technol Assess* 2007;11(49):iii-iv, ix-115. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Shavelle DM, Budoff MJ, LaMont DH,, et al. Exercise testing and electron beam computed tomography in the evaluation of coronary artery disease. *J Am Coll Cardiol* 2000;36(1):32-8. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Shaw LJ, Bairey Merz CN, Pepine CJ,, et al. Insights from the NHLBI-Sponsored Women's Ischemia Syndrome Evaluation (WISE) Study: Part I: gender differences in traditional and novel risk factors, symptom evaluation, and gender-optimized diagnostic strategies. *J Am Coll Cardiol* 2006;47(3 Suppl):S4-S20. *Full-text exclusion reason(s): Not a clinical study report.*



Shaw LJ, Berman DS, Hendel RC, et al. Prognosis by coronary computed tomographic angiography: matched comparison with myocardial perfusion single-photon emission computed tomography. *J Cardiovasc Comput Tomogr* 2008;2(2):93-101. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Shaw LJ, Gillam L, Feinstein S, et al. Use of an intravenous contrast agent (Optison) to enhance echocardiography: efficacy and cost implications. Optison Multicenter Study Group. *Am J Manag Care* 1998;4 Spec No:SP169-76. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Shaw LJ, Hachamovitch R, Berman DS, et al. The economic consequences of available diagnostic and prognostic strategies for the evaluation of stable angina patients: an observational assessment of the value of precatheterization ischemia. Economics of Noninvasive Diagnosis (END) Multicenter Study Group. *J Am Coll Cardiol* 1999;33(3):661-9. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; No outcomes of interest.*

Shaw LJ, Hachamovitch R, Redberg RF. Current evidence on diagnostic testing in women with suspected coronary artery disease: choosing the appropriate test. *Cardiol Rev* 2000;8(1):65-74. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Shaw LJ, Hendel R, Borges-Neto S, et al. Prognostic value of normal exercise and adenosine (99m)Tc-tetrofosmin SPECT imaging: results from the multicenter registry of 4,728 patients. *J Nucl Med* 2003;44(2):134-9. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization.*

Shaw LJ, Hendel RC, Cerquiera M, et al. Ethnic differences in the prognostic value of stress technetium-99m tetrofosmin gated single-photon emission computed tomography myocardial perfusion imaging. *J Am Coll Cardiol* 2005;45(9):1494-504. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Shaw LJ, Marwick TH, Berman DS, et al. Incremental cost-effectiveness of exercise echocardiography vs. SPECT imaging for the evaluation of stable chest pain. *Eur Heart J* 2006;27(20):2448-58. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Shaw LJ, Mieres JH. The role of noninvasive testing in the diagnosis and prognosis of women with suspected CAD. *J Fam Pract* 2005;Suppl:4-5, 7. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Shaw LJ, Min JK, Narula J, et al. Sex differences in mortality associated with computed tomographic angiographic measurements of obstructive and nonobstructive coronary artery disease: an exploratory analysis. *Circ Cardiovasc Imaging* 2010;3(4):473-81. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Shaw LJ, Vasey C, Sawada S, et al. Impact of gender on risk stratification by exercise and dobutamine stress echocardiography: long-term mortality in 4234 women and 6898 men. *Eur Heart J* 2005;26(5):447-56. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization.*

Shaw LJ, Wilson PW, Hachamovitch R, et al. Improved near-term coronary artery disease risk classification with gated stress myocardial perfusion SPECT. *JACC Cardiovasc Imaging* 2010;3(11):1139-48. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Sheikh M, Ben-Nakhi A, Shukkur AM, et al. Accuracy of 64-multidetector-row computed tomography in the diagnosis of coronary artery disease. *Med Princ Pract* 2009;18(4):323-8. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Sheldon SH, Askew JW, 3rd, Klarich KW, et al. Occurrence of atrial fibrillation during dobutamine stress echocardiography: incidence, risk factors, and outcomes. *J Am Soc Echocardiogr* 2011;24(1):86-90. *Full-text*

*exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup; No outcomes of interest.*

Shelley S, Sathyamurthy I, Madhavan,, et al. Adenosine myocardial SPECT—its efficacy and safety and correlation with coronary angiogram. *J Assoc Physicians India* 2003;51:557-60. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Shen Y, Qian JY, Wang MH,, et al. Quantitative and qualitative assessment of non-obstructive left main coronary artery plaques using 64-multislice computed tomography compared with intravascular ultrasound. *Chin Med J (Engl)* 2010;123(7):827-33. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Shenoy MM, Sengupta R, Khanna A. Atrial pacing stress echocardiography: an alternative diagnostic test for chest pain in the elderly. *Am J Geriatr Cardiol* 2002;11(6):404-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Sheth T, Amlani S, Ellins ML,, et al. Computed tomographic coronary angiographic assessment of high-risk coronary anatomy in patients with suspected coronary artery disease and intermediate pretest probability. *Am Heart J* 2008;155(5):918-23. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Sheth TN, Rieber J, Mooyaart EA,, et al. Usefulness of coronary computed tomographic angiography to assess suitability for revascularization in patients with significant coronary artery disease and angina pectoris. *Am J Cardiol* 2006;98(9):1198-201. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Shi H, Aschoff AJ, Brambs HJ,, et al. Multislice CT imaging of anomalous coronary arteries. *Eur Radiol* 2004;14(12):2172-81. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Shi H, Han P, Kong X,, et al. Noninvasive detection of coronary artery stenosis using 16-slice spiral CT: a comparison with selective X-ray coronary angiography. *J Huazhong Univ Sci Technolog Med Sci* 2006;26(3):338-40. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Shiba C, Chikamori T, Hida S,, et al. Important parameters in the detection of left main trunk disease using stress myocardial perfusion imaging. *J Cardiol* 2009;53(1):43-52. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Shimoni S, Goland S, Livshitz S,, et al. Accuracy and long-term prognostic value of pacing stress echocardiography compared with dipyridamole Tl emission computed tomography in patients with a permanent pacemaker and known or suspected coronary artery disease. *Cardiology* 2010;116(3):229-236. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Shimoni S, Zoghbi WA, Xie F,, et al. Real-time assessment of myocardial perfusion and wall motion during bicycle and treadmill exercise echocardiography: comparison with single photon emission computed tomography. *J Am Coll Cardiol* 2001;37(3):741-7. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Shin JH, Pokharna HK, Williams KA,, et al. SPECT myocardial perfusion imaging with prone-only acquisitions: correlation with coronary angiography. *J Nucl Cardiol* 2009;16(4):590-6. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Shirai N, Yamagishi H, Yoshiyama M,, et al. Incremental value of assessment of regional wall motion for detection of multivessel coronary artery disease in exercise (201)Tl gated myocardial perfusion imaging. *J Nucl Med* 2002;43(4):443-50. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Shiyovich A, Jafari J, Blaer Y,, et al. Respiratory stress response: a novel diagnostic method for detection of significant coronary artery disease from finger pulse wave analysis during brief respiratory exercise. *Am J Med Sci* 2010;339(5):440-7. *Full-text exclusion reason(s): Article retracted.*

Shry EA, Eckart RE, Furgerson JL,, et al. Addition of right-sided and posterior precordial leads during stress testing. *Am Heart J* 2003;146(6):1090-4. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Shuman WP, Branch KR, May JM,, et al. Whole-chest 64-MDCT of emergency department patients with nonspecific chest pain: Radiation dose and coronary artery image quality with prospective ECG triggering versus retrospective ECG gating. *AJR Am J Roentgenol* 2009;192(6):1662-7. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Shuman WP, May JM, Branch KR,, et al. Negative ECG-gated cardiac CT in patients with low-to-moderate risk chest pain in the emergency department: 1-year follow-up. *AJR Am J Roentgenol* 2010;195(4):923-7. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Sicari R, Pasanisi E, Venneri L,, et al. Stress echo results predict mortality: a large-scale multicenter prospective international study. *J Am Coll Cardiol* 2003;41(4):589-95. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization.*

Sidhu M, Bhalla M, Chockalingam A,, et al. Pertinence of myocardial perfusion imaging in octogenarians. *Angiology* 2010;61(3):294-6. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Siebelink HM, Blanksma PK, Crijns HJ,, et al. No difference in cardiac event-free survival between positron emission tomography-guided and single-photon emission computed tomography-guided patient management: a prospective, randomized comparison of patients with suspicion of jeopardized myocardium. *J Am Coll Cardiol* 2001;37(1):81-8. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Siennicki J, Kusmierk J, Kovacevic-Kusmierk K,, et al. The effect of image translation table on diagnostic efficacy of myocardial perfusion SPECT studies. *Nucl Med Rev* 2010;13(2):64-69. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Sikdar S, Beach KW, Goldberg SL,, et al. Ultrasonic Doppler vibrometry: measurement of left ventricular wall vibrations associated with coronary artery disease. *Conf Proc IEEE Eng Med Biol Soc* 2006;1:863-6. *Full-text exclusion reason(s): No data for NITs of interest (ECG, ECHO, SPECT, PET, CMR, CTA).*

Sikdar S, Lee JC, Remington J,, et al. Ultrasonic Doppler vibrometry: novel method for detection of left ventricular wall vibrations caused by poststenotic coronary flow. *J Am Soc Echocardiogr* 2007;20(12):1386-92. *Full-text exclusion reason(s): No data for NITs of interest (ECG, ECHO, SPECT, PET, CMR, CTA).*

Simone GL, Mullani NA, Page DA,, et al. Utilization statistics and diagnostic accuracy of a nonhospital-based positron emission tomography center for the detection of coronary artery disease using rubidium-82. *Am J Physiol Imaging* 1992;7(3-4):203-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Simonetti I, Rezai K, Rossen JD,, et al. Physiological assessment of sensitivity of noninvasive testing for coronary artery disease. *Circulation* 1991;83(5 SUPPL.). *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Simova I, Katova T, Denchev S,, et al. Flow-mediated dilatation has an additive value to stress ECG for the diagnosis of angiographically significant coronary atherosclerosis. *J Am Soc Hypertens* 2010;4(4):203-8. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Sinha AM, Mahnken AH, Borghans A,, et al. Multidetector-row computed tomography vs. angiography and intravascular ultrasound for the evaluation of the diameter of proximal coronary arteries. *Int J Cardiol* 2006;110(1):40-5. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Sirol M, Sanz J, Henry P,, et al. Evaluation of 64-slice MDCT in the real world of cardiology: a comparison with conventional coronary angiography. *Arch Cardiovasc Dis* 2009;102(5):433-9. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Sitges M, Azqueta M, Pare C,, et al. Dobutamine stress echocardiography and exercise electrocardiography for risk stratification in medically treated unstable angina. *J Am Soc Echocardiogr* 2000;13(12):1084-90. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Slart RH, Zeebregts CJ, Hillege HL,, et al. Myocardial perfusion reserve after a PET-driven revascularization procedure: a strong prognostic factor. *J Nucl Med* 2011;52(6):873-9. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Smart SC, Bhatia A, Hellman R,, et al. Dobutamine-atropine stress echocardiography and dipyridamole sestamibi scintigraphy for the detection of coronary artery disease: limitations and concordance. *J Am Coll Cardiol* 2000;36(4):1265-73. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Smart SC, Knickelbine T, Malik F,, et al. Dobutamine-atropine stress echocardiography for the detection of coronary artery disease in patients with left ventricular hypertrophy. Importance of chamber size and systolic wall stress. *Circulation* 2000;101(3):258-63. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Smart SC, Knickelbine T, Stoiber TR,, et al. Safety and accuracy of dobutamine-atropine stress echocardiography for the detection of residual stenosis of the infarct-related artery and multivessel disease during the first week after acute myocardial infarction. *Circulation* 1997;95(6):1394-401. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Smit MD, Tio RA, Slart RH,, et al. Myocardial perfusion imaging does not adequately assess the risk of coronary artery disease in patients with atrial fibrillation. *Europace* 2010;12(5):643-8. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

So NM, Lam WW, Li D,, et al. Magnetic resonance angiography of coronary arteries with a 3-dimensional magnetization-prepared true fast imaging with steady-state precession sequence compared with conventional coronary angiography. *Am Heart J* 2005;150(3):530-5. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Sochowski RA, Yvorchuk KJ, Yang Y,, et al. Dobutamine and dipyridamole stress echocardiography in patients with a low incidence of severe coronary artery disease. *J Am Soc Echocardiogr* 1995;8(4):482-7. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Solot G, Hermans J, Merlo P,, et al. Correlation of 99Tcm-sestamibi SPECT with coronary angiography in general hospital practice. *Nucl Med Commun* 1993;14(1):23-9. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Soman P, Khattar R, Senior R,, et al. Inotropic stress with arbutamine is superior to vasodilator stress with dipyridamole for the detection of reversible ischemia with Tc-99m sestamibi single-photon emission computed tomography. *J Nucl Cardiol* 1997;4(5):364-71. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Soman P, Parsons A, Lahiri N,, et al. The prognostic value of a normal Tc-99m sestamibi SPECT study in suspected coronary artery disease. *J Nucl Cardiol* 1999;6(3):252-6. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Sommer T, Hackenbroch M, Hofer U,, et al. Coronary MR angiography at 3.0 T versus that at 1.5 T: initial results in patients suspected of having coronary artery disease. *Radiology* 2005;234(3):718-25. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Soofi MA, Khan SA. Dobutamine stress echocardiography as a prognostic tool for future cardiac events. *Ann Saudi Med* 2008;28(5):371-3. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Soon KH, Chaitowitz I, Cox N, et al. Diagnostic accuracy of 16-slice CT coronary angiography in the evaluation of coronary artery disease. *Australas Radiol* 2007;51(4):365-9. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Soon KH, Cox N, Chaitowitz I, et al. Determining the proportion of coronary segments assessable on 16-slice CT coronary angiography: a brief report. *Australas Radiol* 2007;51(2):139-42. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Soon KH, Kelly AM, Cox N, et al. Practicality, safety and accuracy of computed tomography coronary angiography in the evaluation of low TIMI-risk score chest pain patients: a pilot study. *Emerg Med Australas* 2007;19(2):129-35. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Sosnowski M, Pysz P, Gola A, et al. Coronary artery visualization using a 64-row multi-slice computed tomography in unselected patients with definite or suspected coronary artery disease: a comparison with invasive coronary angiography. *Cardiol J* 2009;16(5):413-7. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Sosnowski M, Pysz P, Szymanski L, et al. Negative calcium score and the presence of obstructive coronary lesions in patients with intermediate CAD probability. *Int J Cardiol* 2011;148(1):e16-e18. *Full-text exclusion reason(s): Data for women not reported as a subgroup; No outcomes of interest.*

Southard J, Baker L, Schaefer S. In search of the false-negative exercise treadmill testing evidence-based use of exercise echocardiography. *Clin Cardiol* 2008;31(1):35-40. *Full-text exclusion reason(s): All women in the study are known to have CAD.*

Soylu O, Celik S, Karakus G, et al. Transthoracic Doppler echocardiographic coronary flow imaging in identification of left anterior descending coronary artery stenosis in patients with left bundle branch block. *Echocardiography* 2008;25(10):1065-70. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Sozzi FB, Civaia F, Rossi P, et al. Long-term follow-up of patients with first-time chest pain having 64-slice computed tomography. *Am J Cardiol* 2011;107(4):516-21. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Sozzi FB, Elhendy A, Roelandt JR, et al. Prognostic value of dobutamine stress echocardiography in patients with diabetes. *Diabetes Care* 2003;26(4):1074-8. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Sozzi FB, Poldermans D, Bax JJ, et al. Second harmonic imaging improves sensitivity of dobutamine stress echocardiography for the diagnosis of coronary artery disease. *Am Heart J* 2001;142(1):153-9. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Srivastava AV, Ananthasubramaniam K, Patel SJ, et al. Prognostic implications of negative dobutamine stress echocardiography in African Americans compared to Caucasians. *Cardiovasc Ultrasound* 2008;6:20. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Staniloff HM, Forrester JS, Berman DS, et al. Prediction of death, myocardial infarction, and worsening chest pain using thallium scintigraphy and exercise electrocardiography. *J Nucl Med* 1986;27(12):1842-8. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Starr H, Powers L, Safranek S. Clinical Inquiries: What is the best noninvasive diagnostic test for women with suspected CAD? *J Fam Pract* 2010;59(9):534-5. *Full-text exclusion reason(s): Not a clinical study report.*

Steel K, Broderick R, Gandla V,, et al. Complementary prognostic values of stress myocardial perfusion and late gadolinium enhancement imaging by cardiac magnetic resonance in patients with known or suspected coronary artery disease. *Circulation* 2009;120(14):1390-400. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Stegger L, Lipke CS, Kies P,, et al. Quantification of left ventricular volumes and ejection fraction from gated <sup>99m</sup>Tc-MIBI SPECT: validation of an elastic surface model approach in comparison to cardiac magnetic resonance imaging, 4D-MSPECT and QGS. *Eur J Nucl Med Mol Imaging* 2007;34(6):900-9. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Stein PD, Beemath A, Kayali F,, et al. Multidetector computed tomography for the diagnosis of coronary artery disease: a systematic review. *Am J Med* 2006;119(3):203-16. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Stein PD, Yaekoub AY, Matta F,, et al. 64-slice CT for diagnosis of coronary artery disease: a systematic review. *Am J Med* 2008;121(8):715-25. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Steingart RM, Hodnett P, Musso J,, et al. Exercise myocardial perfusion imaging in elderly patients. *J Nucl Cardiol* 2002;9(6):573-80. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Stewart RE, Schwaiger M, Molina E,, et al. Comparison of rubidium-82 positron emission tomography and thallium-201 SPECT imaging for detection of coronary artery disease. *Am J Cardiol* 1991;67(16):1303-10. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Stolzmann P, Alkadhi H, Scheffel H,, et al. Combining cardiac magnetic resonance and computed tomography coronary calcium scoring: added value for the assessment of morphological coronary disease? *Int J Cardiovasc Imaging* 2010. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Stolzmann P, Donati OF, Scheffel H,, et al. Low-dose CT coronary angiography for the prediction of myocardial ischaemia. *Eur Radiol* 2010;20(1):56-64. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Stolzmann P, Goetti R, Baumueller S,, et al. Prospective and retrospective ECG-gating for CT coronary angiography perform similarly accurate at low heart rates. *Euro J Radiol* 2011;79(1):85-91. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Stolzmann P, Scheffel H, Leschka S,, et al. Influence of calcifications on diagnostic accuracy of coronary CT angiography using prospective ECG triggering. *AJR Am J Roentgenol* 2008;191(6):1684-9. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Stratmann HG, Mark AL, Walter KE,, et al. Prognostic value of atrial pacing and thallium-201 scintigraphy in patients with stable chest pain. *Am J Cardiol* 1989;64(16):985-90. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Stratmann HG, Williams GA, Wittry MD,, et al. Exercise technetium-99m sestamibi tomography for cardiac risk stratification of patients with stable chest pain. *Circulation* 1994;89(2):615-22. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Strobeck JE, Shen JT, Singh B,, et al. Comparison of a two-lead, computerized, resting ECG signal analysis device, the MultiFunction-CardioGram or MCG (a.k.a. 3DMP), to quantitative coronary angiography for the detection of relevant coronary artery stenosis (>70%) - a meta-analysis of all published trials performed and analyzed in the US. *Int J Med Sci* 2009;6(4):143-55. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Sugihara H, Tamaki N, Mitsunami K,, et al. Prognostic value of 1-day stress/rest electrocardiogram-gated single-photon emission computed tomography using Tc-99m-labeled methoxy-isobutyl isonitrile. *Jpn Circ J* 1998;62(6):405-8. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Sumanen M, Jussila M, Mattila K. Exercise treadmill test may predict clinical outcome among working-aged patients suspected of coronary heart disease in general practice. *Scand J Prim Health Care* 2005;23(1):47-51. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Sumanen M, Mattila K. A negative finding in an exercise test is reliable among elderly people: a follow-up study. *Gerontology* 2007;53(3):159-64. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Sun Z, Cao Y. Multislice CT angiography assessment of left coronary artery: Correlation between bifurcation angle and dimensions and development of coronary artery disease. *Euro J Radiol* 2011;79(2):e90-e95. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; No outcomes of interest.*

Sun Z, Dimpudus FJ, Nugroho J, et al. CT virtual intravascular endoscopy assessment of coronary artery plaques: a preliminary study. *Eur J Radiol* 2010;75(1):e112-9. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Sun Z, Lin C, Davidson R, et al. Diagnostic value of 64-slice CT angiography in coronary artery disease: a systematic review. *Eur J Radiol* 2008;67(1):78-84. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Sundram F, Notghi A, Smith NB. Pharmacological stress myocardial perfusion scintigraphy: use of a modified adenosine protocol in patients with asthma. *Nucl Med Commun* 2009;30(3):217-25. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Sunman H, Yorgun H, Canpolat U, et al. Association between family history of premature coronary artery disease and coronary atherosclerotic plaques shown by multidetector computed tomography coronary angiography. *Int J Cardiol* 2011. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Suratkal V, Shirke M, Lele RD. Treadmill ECG test combined with myocardial perfusion imaging for evaluation of coronary artery disease: analysis of 340 cases. *J Assoc Physicians India* 2003;51:561-4. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Suzuki Y, Slomka PJ, Wolak A, et al. Motion-frozen myocardial perfusion SPECT improves detection of coronary artery disease in obese patients. *J Nucl Med* 2008;49(7):1075-9. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Svane B, Bone D, Holmgren A. Coronary angiography and thallium-201 single photon emission computed tomography in single vessel coronary artery disease. *Acta Radiol* 1990;31(3):237-44. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Swailam S, Abdel-Salam Z, Emil S, et al. Multi-slice computed tomography: Can it adequately rule out left main coronary disease in patients with an intermediate probability of coronary artery disease? *Cardiol J* 2010;17(6):594-8. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Sylvén C, Hagerman I, Ylen M, et al. Variance ECG detection of coronary artery disease—a comparison with exercise stress test and myocardial scintigraphy. *Clin Cardiol* 1994;17(3):132-40. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Taggu W, Cooper AF, Patel NR, et al. Clinical Symptoms and Non-Invasive Testing Fail to Predict the Presence of Significant Coronary Artery Disease in Men and Women Undergoing Coronary Angiography. 12th World Congress on Heart Disease - New Trends in Research, Diagnosis and Treatment, Hyatt Regency Vancouver, Vancouver (Canada), 16-19 Jul 2005; 2005.

Takagi T, Takagi A, Yoshikawa J. Detection of coronary artery disease using delayed strain imaging at 5 min after the termination of exercise stress: head to head comparison with conventional treadmill stress echocardiography. *J Cardiol* 2010;55(1):41-8. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Takase B, Nagata M, Kihara T, et al. Whole-heart dipyridamole stress first-pass myocardial perfusion MRI for the detection of coronary artery disease. *Jpn Heart J* 2004;45(3):475-86. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Takeishi Y, Chiba J, Abe S, et al. Noninvasive identification of left main and three-vessel coronary artery disease by thallium-201 single photon emission computed tomography during adenosine infusion. *Ann Nucl Med* 1994;8(1):1-7. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Takeuchi M, Araki M, Nakashima Y, et al. Comparison of dobutamine stress echocardiography and stress thallium-201 single-photon emission computed tomography for detecting coronary artery disease. *J Am Soc Echocardiogr* 1993;6(6):593-602. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Tamaki N, Yonekura Y, Mukai T, et al. Stress thallium-201 transaxial emission computed tomography: quantitative versus qualitative analysis for evaluation of coronary artery disease. *J Am Coll Cardiol* 1984;4(6):1213-21. *Full-text exclusion reason(s): Population does not include women  $\geq$  age 18; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Tamaki N, Yonekura Y, Senda M, et al. Value and limitation of stress thallium-201 single photon emission computed tomography: comparison with nitrogen-13 ammonia positron tomography. *J Nucl Med* 1988;29(7):1181-8. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Tamarappoo B, Dey D, Shmilovich H, et al. Increased pericardial fat volume measured from noncontrast CT predicts myocardial ischemia by SPECT. *JACC Cardiovasc Imaging* 2010;3(11):1104-12. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Tamarappoo BK, Dey D, Nakazato R, et al. Comparison of the extent and severity of myocardial perfusion defects measured by CT coronary angiography and SPECT myocardial perfusion imaging. *JACC Cardiovasc Imaging* 2010;3(10):1010-9. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Tardif JC, Dore A, Chan KL, et al. Economic impact of contrast stress echocardiography on the diagnosis and initial treatment of patients with suspected coronary artery disease. *J Am Soc Echocardiogr* 2002;15(11):1335-45. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Tartagni F, Dondi M, Limonetti P, et al. Dipyridamole technetium-99m-2-methoxy isobutyl isonitrile tomoscintigraphic imaging for identifying diseased coronary vessels: comparison with thallium-201 stress-rest study. *J Nucl Med* 1991;32(3):369-76. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Tashiro K, Tomiguchi S, Shiraishi S, et al. Clinical usefulness of a collimator distance dependent resolution recovery in myocardial perfusion SPECT: a clinical report from a single institute. *Ann Nucl Med* 2011;25(2):133-7. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup; No outcomes of interest.*

Tawa CB, Baker WB, Kleiman NS, et al. Comparison of adenosine echocardiography, with and without isometric handgrip, to exercise echocardiography in the detection of ischemia in patients with coronary artery disease. *J Am Soc Echocardiogr* 1996;9(1):33-43. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Tejani FH, Thompson RC, Iskandrian AE, et al. Effect of caffeine on SPECT myocardial perfusion imaging during regadenoson pharmacologic stress: rationale and design of a prospective, randomized, multicenter study. *J Nucl Cardiol* 2011;18(1):73-81. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup; No outcomes of interest.*



Tepe SM. Imaging of the heart and coronary arteries. *Cardiovasc Intervent Radiol* 2003;26(SUPPL. 1). *Full-text exclusion reason(s): Conference abstract or trial registry posting.*

Teresinska A, Wnuk J, Konieczna S, et al. Verification of the left ventricular ejection fraction from gated myocardial perfusion studies (GSPECT). *Kardiol Pol* 2005;63(5):465-75; discussion 476-7. *Full-text exclusion reason(s): Data for women not reported as a subgroup; No outcomes of interest.*

Thanigaraj S, Nease RF, Jr., Schechtman KB, et al. Use of contrast for image enhancement during stress echocardiography is cost-effective and reduces additional diagnostic testing. *Am J Cardiol* 2001;87(12):1430-2. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Therre T, Ribal JP, Motreff P, et al. Assessment of cardiac risk before aortic reconstruction: noninvasive work-up using clinical examination, exercise testing, and dobutamine stress echocardiography versus routine coronary arteriography. *Ann Vasc Surg* 1999;13(5):501-8. *Full-text exclusion reason(s): Data for women not reported as a subgroup; No outcomes of interest.*

Thiele H, Plein S, Breeuwer M, et al. Color-encoded semiautomatic analysis of multi-slice first-pass magnetic resonance perfusion: comparison to tetrofosmin single photon emission computed tomography perfusion and X-ray angiography. *Int J Cardiovasc Imaging* 2004;20(5):371-84; discussion 385-7. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Thilo C, Schoepf UJ, Gordon L, et al. Integrated assessment of coronary anatomy and myocardial perfusion using a retractable SPECT camera combined with 64-slice CT: Initial experience. *Euro Radiol* 2009;19(4):845-856. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Thomas C, Brodoefel H, Tsiflikas I, et al. Does clinical pretest probability influence image quality and diagnostic accuracy in dual-source coronary CT angiography? *Acad Radiol* 2010;17(2):212-8. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Thomas D, Meyer C, Strach K, et al. Dobutamine stress tagging and gradient-echo imaging for detection of coronary heart disease at 3 T. *Br J Radiol* 2011;84(997):44-50. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Thomas D, Strach K, Meyer C, et al. Combined myocardial stress perfusion imaging and myocardial stress tagging for detection of coronary artery disease at 3 Tesla. *J Cardiovasc Magn Reson* 2008;10:59. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Thomas GS, Miyamoto MI, Morello AP, 3rd, et al. Technetium 99m sestamibi myocardial perfusion imaging predicts clinical outcome in the community outpatient setting. The Nuclear Utility in the Community (NUC) Study. *J Am Coll Cardiol* 2004;43(2):213-23. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Thomas GS, Prill NV, Majmundar H, et al. Treadmill exercise during adenosine infusion is safe, results in fewer adverse reactions, and improves myocardial perfusion image quality. *J Nucl Cardiol* 2000;7(5):439-46. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization.*

Thompson RC, Heller GV, Johnson LL, et al. Value of attenuation correction on ECG-gated SPECT myocardial perfusion imaging related to body mass index. *J Nucl Cardiol* 2005;12(2):195-202. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Thomson HL, Basmadjian AJ, Rainbird AJ, et al. Contrast echocardiography improves the accuracy and reproducibility of left ventricular remodeling measurements: a prospective, randomly assigned, blinded study. *J Am Coll Cardiol* 2001;38(3):867-75. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup; No outcomes of interest.*

- Tian J, Zhang G, Wang X,, et al. Exercise echocardiography: feasibility and value for detection of coronary artery disease. *Chin Med J (Engl)* 1996;109(5):381-4. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*
- Toledo E, Jacobs LD, Lodato JA,, et al. Quantitative diagnosis of stress-induced myocardial ischemia using analysis of contrast echocardiographic parametric perfusion images. *Eur J Echocardiogr* 2006;7(3):217-25. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*
- Tolstrup K, Madsen BE, Ruiz JA,, et al. Non-invasive resting magnetocardiographic imaging for the rapid detection of ischemia in subjects presenting with chest pain. *Cardiology* 2006;106(4):270-6. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*
- Tomaszuk-Kazberuk A, Sobkowicz B, Malyszko J,, et al. Perfusion defects on real-time myocardial contrast echocardiography predict higher mortality in patients with end-stage renal disease - a 3-year follow-up. *Ren Fail* 2010;32(10):1160-6. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*
- Toriyama T, Takase H, Goto T,, et al. Coronary artery disease investigated using 99mTc-tetrofosmin myocardial SPECT. *European journal of clinical investigation*; 2007:478-82.
- Toumanidis ST, Pantelia MI, Trika CO,, et al. Detection of coronary artery disease in the presence of left ventricular atrophy. *Int J Cardiol* 1996;57(3):245-55. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*
- Trabold T, Buchgeister M, Kuttner A,, et al. Estimation of radiation exposure in 16-detector row computed tomography of the heart with retrospective ECG-gating. *Rofo* 2003;175(8):1051-5. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*
- Travin MI, Dessouki A, Cameron T,, et al. Use of exercise technetium-99m sestamibi SPECT imaging to detect residual ischemia and for risk stratification after acute myocardial infarction. *Am J Cardiol* 1995;75(10):665-9. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*
- Travin MI, Laraia PJ. The prognostic value of stress radionuclide myocardial perfusion imaging. *Prim Cardiol* 1994(20):44-52. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*
- Truth MG, Reyes GA, He ZX,, et al. Tolerance and diagnostic accuracy of an abbreviated adenosine infusion for myocardial scintigraphy: a randomized, prospective study. *J Nucl Cardiol* 2001;8(5):548-54. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*
- Tsai IC, Lee T, Lee WL,, et al. Use of 40-detector row computed tomography before catheter coronary angiography to select early conservative versus early invasive treatment for patients with low-risk acute coronary syndrome. *J Comput Assist Tomogr* 2007;31(2):258-64. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*
- Tsai MF, Kao PF, Tzen KY. Improved diagnostic performance of thallium-201 myocardial perfusion scintigraphy in coronary artery disease: from planar to single photon emission computed tomography imaging. *Chang Gung Med J* 2002;25(8):522-30. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*
- Tsiflikas I, Brodoefel H, Reimann AJ,, et al. Coronary CT angiography with dual source computed tomography in 170 patients. *Eur J Radiol* 2010;74(1):161-5. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*
- Tsiflikas I, Drosch T, Brodoefel H,, et al. Diagnostic accuracy and image quality of cardiac dual-source computed tomography in patients with arrhythmia. *Int J Cardiol* 2010;143(1):79-85. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Tsougos E, Panou F, Paraskevaïdis I, et al. Exercise-induced changes in E/E' ratio in patients with suspected coronary artery disease. *Coron Artery Dis* 2008;19(6):405-11. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Tsutsui JM, Elhendy A, Anderson JR, et al. Prognostic value of dobutamine stress myocardial contrast perfusion echocardiography. *Circulation* 2005;112(10):1444-50. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Tsutsui JM, Elhendy A, Xie F, et al. Safety of dobutamine stress real-time myocardial contrast echocardiography. *J Am Coll Cardiol* 2005;45(8):1235-42. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Tsutsui JM, Osorio AF, Lario FA, et al. Comparison of safety and efficacy of the early injection of atropine during dobutamine stress echocardiography with the conventional protocol. *Am J Cardiol* 2004;94(11):1367-72. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Tsutsui JM, Xie F, Cloutier D, et al. Real-time dobutamine stress myocardial perfusion echocardiography predicts outcome in the elderly. *Eur Heart J* 2008;29(3):377-85. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Tsutsui JM, Xie F, McGrain AC, et al. Comparison of low-mechanical index pulse sequence schemes for detecting myocardial perfusion abnormalities during vasodilator stress echocardiography. *Am J Cardiol* 2005;95(5):565-70. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Tsutsui JM, Xie F, O'Leary EL, et al. Diagnostic accuracy and prognostic value of dobutamine stress myocardial contrast echocardiography in patients with suspected acute coronary syndromes. *Echocardiography* 2005;22(6):487-95. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Turgut B, Unlu M, Temiz NH, et al. Dobutamine Tc-99m furifosmin SPECT in detection of coronary artery disease: evaluation of same day, rest-stress protocol. *Ann Nucl Med* 2003;17(7):531-9. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Turkvatan A, Biyikoglu SF, Buyukbayraktar F, et al. Clinical value of 16-slice multidetector computed tomography in symptomatic patients with suspected coronary artery disease. *Acta Radiol* 2008;49(4):400-8. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Turner AS, Nathan MC, Watson OF, et al. The correlation of the computer quantitated treadmill exercise electrocardiogram with cinearteriographic assessment of coronary artery disease. *N Z Med J* 1979;89(630):115-8. *Full-text exclusion reason(s): Population does not include women  $\geq$  age 18; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Turner DA, Battle WE, Deshmukh H, et al. The predictive value of myocardial perfusion scintigraphy after stress in patients without previous myocardial infarction. *J Nucl Med* 1978;19(3):249-55. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Ueno K, Anzai T, Jinzaki M, et al. Increased epicardial fat volume quantified by 64-multidetector computed tomography is associated with coronary atherosclerosis and totally occlusive lesions. *Circ J* 2009;73(10):1927-33. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup; No outcomes of interest.*

Ueno K, Anzai T, Jinzaki M, et al. Diagnostic capacity of 64-slice multidetector computed tomography for acute coronary syndrome in patients presenting with acute chest pain. *Cardiology* 2009;112(3):211-8. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Ugolini P, Pressacco J, Lesperance J, et al. Evaluation of coronary atheroma by 64-slice multidetector computed tomography: Comparison with intravascular ultrasound and angiography. *Can J Cardiol* 2009;25(11):641-7. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Ulimoen GR, Gjonnaess E, Atar D,, et al. Noninvasive coronary angiography with 64-channel multidetector computed tomography in patients with acute coronary syndrome. *Acta Radiol* 2008;49(10):1140-4. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Underwood DA, Rincon G, Yiannikas J,, et al. Clinical usefulness of thallium-201 scintigraphy in the study of coronary artery disease: a comparison of two exercise systems. *Cleve Clin Q* 1982;49(2):73-8. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Underwood SR, Anagnostopoulos C, Cerqueira M,, et al. Myocardial perfusion scintigraphy: the evidence. *Eur J Nucl Med Mol Imaging* 2004;31(2):261-91. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Underwood SR, Godman B, Salyani S,, et al. Economics of myocardial perfusion imaging in Europe—the EMPIRE Study. *Eur Heart J* 1999;20(2):157-66. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Uthamalingam S, Zheng H, Leavitt M,, et al. Exercise-induced ST-segment elevation in ECG lead aVR is a useful indicator of significant left main or ostial LAD coronary artery stenosis. *JACC Cardiovasc Imaging* 2011;4(2):176-86. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Vacanti LJ, Sposito AC, Sespedes L,, et al. In comparison to the myocardial perfusion scintigraphy, a treadmill stress test is a viable, efficient and cost effective option to predict cardiovascular events in elderly patients. *Arq Bras Cardiol* 2007;88(5):531-6. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Valdiviezo C, Motivala AA, Hachamovitch R,, et al. The significance of transient ischemic dilation in the setting of otherwise normal SPECT radionuclide myocardial perfusion images. *J Nucl Cardiol* 2011;18(2):220-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Valeti US, Miller TD, Hodge DO,, et al. Exercise single-photon emission computed tomography provides effective risk stratification of elderly men and elderly women. *Circulation* 2005;111(14):1771-6. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization.*

van der Bijl N, Joemai RM, Geleijns J,, et al. Assessment of Agatston coronary artery calcium score using contrast-enhanced CT coronary angiography. *AJR Am J Roentgenol* 2010;195(6):1299-305. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

van Dijkman PR, Kuijpers DA, Blom BM,, et al. Dobutamine stress magnetic resonance imaging: a valuable method in the noninvasive diagnosis of ischemic heart disease. *J Electrocardiol* 2002;35 Suppl:57-9. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

van Geuns RJ, Oudkerk M, Rensing BJ,, et al. Comparison of coronary imaging between magnetic resonance imaging and electron beam computed tomography. *Am J Cardiol* 2002;90(1):58-63. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

van Geuns RJ, Wielopolski PA, de Bruin HG,, et al. MR coronary angiography with breath-hold targeted volumes: preliminary clinical results. *Radiology* 2000;217(1):270-7. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Van Lingen R, Kakani N, Veitch A,, et al. Prognostic and accuracy data of multidetector CT coronary angiography in an established clinical service. *Clin Radiol* 2009;64(6):601-7. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Van Train KF, Areeda J, Garcia EV,, et al. Quantitative same-day rest-stress technetium-99m-sestamibi SPECT: definition and validation of stress normal limits and criteria for abnormality. *J Nucl Med* 1993;34(9):1494-502. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Van Train KF, Maddahi J, Berman DS,, et al. Quantitative analysis of tomographic stress thallium-201 myocardial scintigrams: a multicenter trial. *J Nucl Med* 1990;31(7):1168-79. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Van Velzen JE, Schuijf JD, De Graaf FR,, et al. Diagnostic performance of non-invasive multidetector computed tomography coronary angiography to detect coronary artery disease using different endpoints: Detection of significant stenosis vs. detection of atherosclerosis. *Euro Heart J* 2011;32(5):637-645. *Full-text exclusion reason(s): Data for women not reported as a subgroup; No outcomes of interest.*

Van Werkhoven JM, Cademartiri F, Seitun S,, et al. Diabetes: prognostic value of CT coronary angiography—comparison with a nondiabetic population. *Radiology* 2010;256(1):83-92. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

van Werkhoven JM, de Boer SM, Schuijf JD,, et al. Impact of clinical presentation and pretest likelihood on the relation between calcium score and computed tomographic coronary angiography. *Am J Cardiol* 2010;106(12):1675-9. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

van Werkhoven JM, Gaemperli O, Schuijf JD,, et al. Multislice computed tomography coronary angiography for risk stratification in patients with an intermediate pretest likelihood. *Heart* 2009;95(19):1607-11. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

van Werkhoven JM, Heijenbrok MW, Schuijf JD,, et al. Diagnostic accuracy of 64-slice multislice computed tomographic coronary angiography in patients with an intermediate pretest likelihood for coronary artery disease. *Am J Cardiol* 2010;105(3):302-5. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

van Werkhoven JM, Heijenbrok MW, Schuijf JD,, et al. Combined non-invasive anatomical and functional assessment with MSCT and MRI for the detection of significant coronary artery disease in patients with an intermediate pre-test likelihood. *Heart* 2010;96(6):425-31. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

van Werkhoven JM, Schuijf JD, Gaemperli O,, et al. Prognostic value of multislice computed tomography and gated single-photon emission computed tomography in patients with suspected coronary artery disease. *J Am Coll Cardiol* 2009;53(7):623-32. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

van Werkhoven JM, Schuijf JD, Gaemperli O,, et al. Incremental prognostic value of multi-slice computed tomography coronary angiography over coronary artery calcium scoring in patients with suspected coronary artery disease. *Eur Heart J* 2009;30(21):2622-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

van Werkhoven JM, Schuijf JD, Jukema JW,, et al. Anatomic correlates of a normal perfusion scan using 64-slice computed tomographic coronary angiography. *Am J Cardiol* 2008;101(1):40-5. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Vanhoenacker PK, Decramer I, Bladt O,, et al. Detection of non-ST-elevation myocardial infarction and unstable angina in the acute setting: meta-analysis of diagnostic performance of multi-detector computed tomographic angiography. *BMC Cardiovasc Disord* 2007;7:39. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Vanhoenacker PK, Heijenbrok-Kal MH, Van Heste R,, et al. Diagnostic performance of multidetector CT angiography for assessment of coronary artery disease: meta-analysis. *Radiology* 2007;244(2):419-28. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Vanzetto G, Ormezzano O, Fagret D,, et al. Long-term additive prognostic value of thallium-201 myocardial perfusion imaging over clinical and exercise stress test in low to intermediate risk patients : study in 1137 patients with 6-year follow-up. *Circulation* 1999;100(14):1521-7. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Vavere AL, Arbab-Zadeh A, Rochitte CE,, et al. Coronary artery stenoses: accuracy of 64-detector row CT angiography in segments with mild, moderate, or severe calcification—A subanalysis of the CORE-64 trial. *Radiology* 2011. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Vegsundvag J, Holte E, Wiseth R,, et al. Coronary flow velocity reserve in the three main coronary arteries assessed with transthoracic doppler: A comparative study with quantitative coronary angiography. *Journal of the American*

Society of Echocardiography 2011;24(7):758-767. *Full-text exclusion reason(s): No data for NITs of interest (ECG, ECHO, SPECT, PET, CMR, CTA); Data for women not reported as a subgroup.*

Vegsundvag J, Holte E, Wiseth R,, et al. Transthoracic echocardiography for imaging of the different coronary artery segments: a feasibility study. *Cardiovasc Ultrasound* 2009;7:58. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup; No outcomes of interest.*

Venkatesh V, Ellins ML, Yang S,, et al. Incremental detection of coronary artery disease by assessment of non-calcified plaque on coronary CT angiography. *Clin Radiol* 2009;64(3):250-5. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Verani MS, Mahmarian JJ, Hixson JB,, et al. Diagnosis of coronary artery disease by controlled coronary vasodilation with adenosine and thallium-201 scintigraphy in patients unable to exercise. *Circulation* 1990;82(1):80-7. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup; No outcomes of interest.*

Verani MS, Marcus ML, Razzak MA,, et al. Sensitivity and specificity of thallium-201 perfusion scintigrams under exercise in the diagnosis of coronary artery disease. *J Nucl Med* 1978;19(7):773-82. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Verna E, Ceriani L, Giovannella L,, et al. "False-positive" myocardial perfusion scintigraphy findings in patients with angiographically normal coronary arteries: insights from intravascular sonography studies. *J Nucl Med* 2000;41(12):1935-40. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Vigna C, Stanislaw M, De Rito V,, et al. Dipyridamole stress echocardiography vs dipyridamole sestamibi scintigraphy for diagnosing coronary artery disease in left bundle-branch block. *Chest* 2001;120(5):1534-9. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Visser CA, van der Wieken RL, Kan G,, et al. Comparison of two-dimensional echocardiography with radionuclide angiography during dynamic exercise for the detection of coronary artery disease. *Am Heart J* 1983;106(3):528-34. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Vitarelli A, Conde Y, Luzzi MF,, et al. Transesophageal dobutamine stress echocardiography with tissue Doppler imaging for detection and assessment of coronary artery disease. *J Investig Med* 2001;49(6):534-43. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Vitarelli A, Luzzi MF, Penco M,, et al. On-line quantitative assessment of left ventricular filling during dobutamine stress echocardiography: a useful addition to conventional wall motion scoring. *Int J Cardiol* 1997;59(1):57-69. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Vivekananthan DP, Blackstone EH, Pothier CE,, et al. Heart rate recovery after exercise is a predictor of mortality, independent of the angiographic severity of coronary disease. *J Am Coll Cardiol* 2003;42(5):831-8. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup; No outcomes of interest.*

Vogel R, Indermuhle A, Meier P,, et al. Quantitative stress echocardiography in coronary artery disease using contrast-based myocardial blood flow measurements: prospective comparison with coronary angiography. *Heart* 2009;95(5):377-84. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Vogel-Claussen J, Skrok J, Dombroski D,, et al. Comprehensive adenosine stress perfusion MRI defines the etiology of chest pain in the emergency room: Comparison with nuclear stress test. *J Magn Reson Imaging* 2009;30(4):753-62. *Full-text exclusion reason(s): No outcomes of interest.*

Voros S, Rinehart S, Qian Z., et al. Prospective validation of standardized, 3-dimensional, quantitative coronary computed tomographic plaque measurements using radiofrequency backscatter intravascular ultrasound as reference standard in intermediate coronary arterial lesions: results from the ATLANTA (assessment of tissue characteristics, lesion morphology, and hemodynamics by angiography with fractional flow reserve, intravascular ultrasound and virtual histology, and noninvasive computed tomography in atherosclerotic plaques) I study. *JACC Cardiovasc Interv* 2011;4(2):198-208. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Vural M, Ucar O, Selvi NA., et al. Assessment of global left ventricular systolic function with multidetector. *Diagn Interv Radiol* 2010;16(3):236-240. *Full-text exclusion reason(s): Data for women not reported as a subgroup; No outcomes of interest.*

Wagdi P, Alkadhi H. The impact of cardiac CT on the appropriate utilization of catheter coronary angiography. *Int J Cardiovasc Imaging* 2010;26(3):333-44. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup; No outcomes of interest.*

Wagdi P, Kaufmann U, Fluri M., et al. High dose dipyridamole as a pharmacological stress test during cardiac catheterisation in patients with coronary artery disease. *Heart* 1996;75(3):247-51. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Wagner M, Rosler R, Lembcke A., et al. Whole-heart coronary magnetic resonance angiography at 1.5 Tesla: does a blood-pool contrast agent improve diagnostic accuracy? *Invest Radiol* 2011;46(3):152-9. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Wagner S, Schuster S, Zahn R., et al. Postinfarction stress testing and one year outcome of stable patients after myocardial infarction treated with thrombolytics. *Eur J Med Res* 1996;1(12):575-81. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Wahl A, Paetsch I, Gollesch A., et al. Safety and feasibility of high-dose dobutamine-atropine stress cardiovascular magnetic resonance for diagnosis of myocardial ischaemia: experience in 1000 consecutive cases. *Eur Heart J* 2004;25(14):1230-6. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Wainwright RJ, Maisey MN, Sowton E. Segmental quantitative analysis of digital thallium-201 myocardial scintigrams in diagnosis of coronary artery disease. Comparison with rest and exercise electrocardiography and coronary arteriography. *Br Heart J* 1981;46(5):478-85. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Wakasugi S, Shibata N, Kobayashi T., et al. Specific perfusion pattern in stress 201Tl myocardial scintigraphy of left main coronary artery disease. *Eur J Nucl Med* 1986;12(8):369-74. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Walimbe V, Jaber WA, Garcia MJ., et al. Multimodality cardiac stress testing: combining real-time 3-dimensional echocardiography and myocardial perfusion SPECT. *J Nucl Med* 2009;50(2):226-30. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Wallace EL, Morgan TM, Walsh TF., et al. Dobutamine cardiac magnetic resonance results predict cardiac prognosis in women with known or suspected ischemic heart disease. *JACC Cardiovasc Imaging* 2009;2(3):299-307. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Study does not compare one NIT to another or to diagnostic cardiac catheterization.*

Wallhaus TR, Lacy J, Stewart R., et al. Copper-62-pyruvaldehyde bis(N-methyl-thiosemicarbazone) PET imaging in the detection of coronary artery disease in humans. *J Nucl Cardiol* 2001;8(1):67-74. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Walsh TF, Dall'Armellina E, Chughtai H., et al. Adverse effect of increased left ventricular wall thickness on five year outcomes of patients with negative dobutamine stress. *J Cardiovasc Magn Reson* 2009;11:25. *Full-text*

*exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Wang Q, Qin J, Gai LY, et al. A pilot study on diagnosis of coronary artery disease using computed tomography first-pass myocardial perfusion imaging at rest. *Journal of Zhejiang University: Science B* 2011;12(6):485-491. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Warner MF, Pippin JJ, DiSciascio G, et al. Assessment of thallium scintigraphy and echocardiography during dobutamine infusion for the detection of coronary artery disease. *Cathet Cardiovasc Diagn* 1993;29(2):122-7. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Watanabe H, Hozumi T, Hirata K, et al. Noninvasive coronary flow velocity reserve measurement in the posterior descending coronary artery for detecting coronary stenosis in the right coronary artery using contrast-enhanced transthoracic Doppler echocardiography. *Echocardiography* 2004;21(3):225-33. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Watanabe K, Sekiya M, Ikeda S, et al. Comparison of adenosine triphosphate and dipyridamole in diagnosis by thallium-201 myocardial scintigraphy. *J Nucl Med* 1997;38(4):577-81. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Watkins MW, Hesse B, Green CE, et al. Detection of coronary artery stenosis using 40-channel computed tomography with multi-segment reconstruction. *Am J Cardiol* 2007;99(2):175-81. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Watkins S, McGeoch R, Lyne J, et al. Validation of magnetic resonance myocardial perfusion imaging with fractional flow reserve for the detection of significant coronary heart disease. *Circulation* 2009;120(22):2207-13. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Wehrschoetz M, Wehrschoetz E, Schuchlenz H, et al. Accuracy of MSCT coronary angiography with 64 row CT scanner—Facing the facts. *Clinical Medicine Insights: Cardiology* 2010;4:15-22. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Wei K, Crouse L, Weiss J, et al. Comparison of usefulness of dipyridamole stress myocardial contrast echocardiography to technetium-99m sestamibi single-photon emission computed tomography for detection of coronary artery disease (PB127 Multicenter Phase 2 Trial results). *Am J Cardiol* 2003;91(11):1293-8. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Wei L, Kadoya M, Momose M, et al. Serial assessment of left ventricular function in various patient groups with Tl-201 gated myocardial perfusion SPECT. *Radiat Med* 2007;25(2):65-72. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Weigold WG, Abbara S, Achenbach S, et al. Standardized medical terminology for cardiac computed tomography: a report of the Society of Cardiovascular Computed Tomography. *J Cardiovasc Comput Tomogr* 2011;5(3):136-44. *Full-text exclusion reason(s): Not a clinical study report.*

Weininger M, Schoepf UJ, Ramachandra A, et al. Adenosine-stress dynamic real-time myocardial perfusion CT and adenosine-stress first-pass dual-energy myocardial perfusion CT for the assessment of acute chest pain: Initial results. *Eur J Radiol* 2010. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup; No outcomes of interest.*

Weinsaft JW, Gade CL, Wong FJ, et al. Diagnostic impact of SPECT image display on assessment of obstructive coronary artery disease. *J Nucl Cardiol* 2007;14(5):659-68. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Weinsaft JW, Manoushagian SJ, Patel T, et al. Stress-induced ST-segment deviation in relation to the presence and severity of coronary artery disease in patients with normal myocardial perfusion imaging. *Coron Artery Dis* 2009;20(1):41-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*



Weiss MB, Narasimhadevara SM, Feng GQ, et al. Computer-enhanced frequency-domain and 12-lead electrocardiography accurately detect abnormalities consistent with obstructive and nonobstructive coronary artery disease. *Heart Dis* 2002;4(1):2-12. *Full-text exclusion reason(s): No data for NITs of interest (ECG, ECHO, SPECT, PET, CMR, CTA).*

Weiss SJ, Ernst AA, Godorov G, et al. Bioimpedance-derived differences in cardiac physiology during exercise stress testing in low-risk chest pain patients. *South Med J* 2003;96(11):1121-7. *Full-text exclusion reason(s): Data for women not reported as a subgroup; No outcomes of interest.*

Wenger NK. Coronary heart disease in women: highlights of the past 2 years—stepping stones, milestones and obstructing boulders. *Nat Clin Pract Cardiovasc Med* 2006;3(4):194-202. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Wenger NK, Shaw LJ, Vaccarino V. Coronary heart disease in women: update 2008. *Clin Pharmacol Ther* 2008;83(1):37-51. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Wertman BM, Cheng VY, Kar S, et al. Characterization of complex coronary artery stenosis morphology by coronary computed tomographic angiography. *JACC Cardiovasc Imaging* 2009;2(8):950-8. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Weustink AC, Neeffjes LA, Kyrzopoulos S, et al. Impact of heart rate frequency and variability on radiation exposure, image quality, and diagnostic performance in dual-source spiral CT coronary angiography. *Radiology* 2009;253(3):672-80. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Weustink AC, Neeffjes LA, Rossi A, et al. Diagnostic performance of exercise bicycle testing and single-photon emission computed tomography: comparison with 64-slice computed tomography coronary angiography. *Int J Cardiovasc Imaging* 2011. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Wexler O, Yoder SR, Elder JL, et al. Effect of gender on cardiovascular risk stratification with ECG gated SPECT left ventricular volume indices and ejection fraction. *J Nucl Cardiol* 2009;16(1):28-37. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization.*

White CS, Kuo D, Kelemen M, et al. Chest pain evaluation in the emergency department: can MDCT provide a comprehensive evaluation? *AJR Am J Roentgenol* 2005;185(2):533-40. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Wijesekera NT, Padley SPG, Ansede G, et al. Is there a role for 64-multi-detector CT coronary angiography in octogenarians? A single-centre experience. *Br J Cardiol* 2010;17(1):40-43. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Williams BR, Mullani NA, Jansen DE, et al. A retrospective study of the diagnostic accuracy of a community hospital-based PET center for the detection of coronary artery disease using rubidium-82. *J Nucl Med* 1994;35(10):1586-92. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Williams KA, Schneider CM. Increased stress right ventricular activity on dual isotope perfusion SPECT: a sign of multivessel and/or left main coronary artery disease. *J Am Coll Cardiol* 1999;34(2):420-7. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Winter R, Gudmundsson P, Willenheimer R. Real-time perfusion adenosine stress echocardiography in the coronary care unit: a feasible bedside tool for predicting coronary artery stenosis in patients with acute coronary syndrome. *Eur J Echocardiogr* 2005;6(1):31-40. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Wittlinger T, Voigtlander T, Rohr M, et al. Magnetic resonance imaging of coronary artery occlusions in the navigator technique. *Int J Cardiovasc Imaging* 2002;18(3):203-11; discussion 213-5. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Wolak A, Gutstein A, Cheng VY,, et al. Dual-source coronary computed tomography angiography in patients with atrial fibrillation: initial experience. *J Cardiovasc Comput Tomogr* 2008;2(3):172-80. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Wolber T, Maeder M, Weilenmann D,, et al. Integration of B-type natriuretic peptide levels with clinical data and exercise testing for predicting coronary artery disease. *Am J Cardiol* 2006;98(6):764-7. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Wolff SD, Schwitter J, Coulden R,, et al. Myocardial first-pass perfusion magnetic resonance imaging: a multicenter dose-ranging study. *Circulation* 2004;110(6):732-7. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Wong RC, Sinha AK, Mahadevan M,, et al. Diagnostic utility, safety, and cost-effectiveness of emergency department-initiated early scheduled technetium-99m single photon emission computed tomography imaging followed by expedited outpatient cardiac clinic visits in acute chest pain syndromes. *Emergency Radiology* 2010;17(5):375-380. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Wright DJ, Williams SG, Lindsay HS,, et al. Assessment of adenosine, arbutamine and dobutamine as pharmacological stress agents during (99m)Tc-tetrofosmin SPECT imaging: a randomized study. *Nucl Med Commun* 2001;22(12):1305-11. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Wu CC, Ho YL, Kao SL,, et al. Dobutamine stress echocardiography for detecting coronary artery disease. *Cardiology* 1996;87(3):244-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Wu MC, Chin KC, Lin KH,, et al. Diagnostic efficacy of a low-dose 32-projection SPECT 99mTc-sestamibi myocardial perfusion imaging protocol in routine practice. *Nucl Med Commun* 2009;30(2):140-7. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Wu YW, Huang PJ, Su MY,, et al. Myocardium viability assessed by delayed contrast-enhanced magnetic resonance imaging in patients with severe ischemic heart failure: A comparison with thallium SPECT and dobutamine echocardiography. *World Heart Journal* 2008;1(1):57-68. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Xiao Y, Tian JM, Gong J,, et al. [Screening for consecutive patient population with suspected CAD by 64-slice CT coronary angiography]. *Zhonghua Yu Fang Yi Xue Za Zhi* 2007;41 Suppl:134-7. *Full-text exclusion reason(s): Non-English.*

Xie F, Hankins J, Mahrous HA,, et al. Detection of coronary artery disease with a continuous infusion of definity ultrasound contrast during adenosine stress real time perfusion echocardiography. *Echocardiography* 2007;24(10):1044-50. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Xie F, Tsutsui JM, McGrain AC,, et al. Comparison of dobutamine stress echocardiography with and without real-time perfusion imaging for detection of coronary artery disease. *Am J Cardiol* 2005;96(4):506-11. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Xu L, Yang L, Fan Z,, et al. Diagnostic performance of 320-detector CT coronary angiography in patients with atrial fibrillation: preliminary results. *Eur Radiol* 2011;21(5):936-43. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup; No outcomes of interest.*

Xu Y, Fish M, Gerlach J,, et al. Combined quantitative analysis of attenuation corrected and non-corrected myocardial perfusion SPECT: Method development and clinical validation. *J Nucl Cardiol* 2010;17(4):591-599. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Xu Y, Tang L, Zhu X,, et al. Comparison of dual-source CT coronary angiography and conventional coronary angiography for detecting coronary artery disease. *Int J Cardiovasc Imaging* 2010;26 Suppl 1:75-81. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Yalcin H, Aktas A, Erol T,, et al. Myocardial perfusion SPECT and dobutamine stress tissue Doppler imaging in evaluation of patients with stable angina pectoris. *Anadolu Kardiyol Derg* 2010;10(4):334-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Yamabe H, Fujiwara S, Rin K,, et al. Resting 123I-BMIPP scintigraphy for detection of organic coronary stenosis and therapeutic outcome in patients with chest pain. *Ann Nucl Med* 2000;14(3):187-92. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Yamada AT, Soares J, Jr., Meneghetti JC,, et al. Planar myocardial perfusion imaging for evaluation of patients with acute chest pain. *Int J Cardiol* 2004;97(3):447-53. *Full-text exclusion reason(s): All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Yamada K, Hirai M, Abe K,, et al. Diagnostic usefulness of postexercise systolic blood pressure response for detection of coronary artery disease in patients with echocardiographic left ventricular hypertrophy. *Can J Cardiol* 2004;20(7):705-11. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Yamada T, Sawada T, Yamano T,, et al. Evaluation of coronary arterial stenoses using 2D magnetic resonance coronary angiography. *Minimally Invasive Therapy and Allied Technologies* 2002;11(1):7-15. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Yamaguchi M, Shimizu M, Ino H,, et al. Diagnostic usefulness of the post-exercise systolic blood pressure response for the detection of coronary artery disease in patients with diabetes mellitus. *Jpn Circ J* 2000;64(12):949-52. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Yang CW, Carr JC, Francois CJ,, et al. Coronary magnetic resonance angiography using magnetization-prepared contrast-enhanced breath-hold volume-targeted imaging (MPCE-VCATS). *Invest Radiol* 2006;41(8):639-44. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Yang L, Zhang Z, Fan Z,, et al. 64-MDCT coronary angiography of patients with atrial fibrillation: influence of heart rate on image quality and efficacy in evaluation of coronary artery disease. *AJR Am J Roentgenol* 2009;193(3):795-801. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Yang MF, Dou KF, Liu XJ,, et al. Prognostic value of normal exercise 99mTc-sestamibi myocardial tomography in patients with angiographic coronary artery disease. *Nucl Med Commun* 2006;27(4):333-8. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Yang PC, Santos JM, Nguyen PK,, et al. Dynamic real-time architecture in magnetic resonance coronary angiography—a prospective clinical trial. *J Cardiovasc Magn Reson* 2004;6(4):885-94. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Yang Q, Li K, Liu X,, et al. Contrast-enhanced whole-heart coronary magnetic resonance angiography at 3.0-T: a comparative study with X-ray angiography in a single center. *J Am Coll Cardiol* 2009;54(1):69-76. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Yang WI, Hur J, Ko YG,, et al. Assessment of tissue characteristics of noncalcified coronary plaques by 64-slice computed tomography in comparison with integrated backscatter intravascular ultrasound. *Coron Artery Dis* 2010;21(3):168-74. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Yang X, Gai LY, Li P,, et al. Diagnostic accuracy of dual-source CT angiography and coronary risk stratification. *Vasc Health Risk Manag* 2010;6:935-41. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Yanik A, Yetkin E, Senen K., et al. Value of dobutamine stress echocardiography for diagnosis of coronary artery disease in patients with left bundle branch blockage. *Coron Artery Dis* 2000;11(7):545-8. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Yao SS, Bangalore S, Chaudhry FA. Prognostic implications of stress echocardiography and impact on patient outcomes: An effective gatekeeper for coronary angiography and revascularization. *Journal of the American Society of Echocardiography* 2010;23(8):832-839. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Yao SS, Qureshi E, Sherrid MV., et al. Practical applications in stress echocardiography: risk stratification and prognosis in patients with known or suspected ischemic heart disease. *J Am Coll Cardiol* 2003;42(6):1084-90. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Yao SS, Shah A, Bangalore S., et al. Transient ischemic left ventricular cavity dilation is a significant predictor of severe and extensive coronary artery disease and adverse outcome in patients undergoing stress echocardiography. *J Am Soc Echocardiogr* 2007;20(4):352-8. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Yao SS, Weinberg C, Bangalore S., et al. The changing paradigm of stress echocardiography: risk stratification, prognosis, and future directions. *Hosp Pract (Minneapolis)* 2010;38(3):26-39. *Full-text exclusion reason(s): Not original peer-reviewed data or a secondary analysis of RCT, prospective or retrospective observational study, or registry.*

Yao Z, Liu XJ, Shi R., et al. A comparison of 99mTc-MIBI myocardial SPET with electron beam computed tomography in the assessment of coronary artery disease. *Eur J Nucl Med* 1997;24(9):1115-20. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Yao Z, Liu XJ, Shi RF., et al. A comparison of 99Tcm-MIBI myocardial SPET and electron beam computed tomography in the assessment of coronary artery disease in two different age groups. *Nucl Med Commun* 2000;21(1):43-8. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Yao ZM, Li W, Qu WY., et al. Comparison of 9(9m)Tc-methoxyisobutylisonitrile myocardial single-photon emission computed tomography and electron beam computed tomography for detecting coronary artery disease in patients with no myocardial infarction. *Chin Med J (Engl)* 2004;117(5):700-705. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Yap LB, Arshad W, Jain A., et al. Significance of ST depression during exercise treadmill stress and adenosine infusion myocardial perfusion imaging. *Int J Cardiovasc Imaging* 2005;21(2-3):253-8; discussion 259-60. *Full-text exclusion reason(s): All women in the study are known to have CAD.*

Yeo TC, Ling LH, Ng WL., et al. Dobutamine stress echocardiography: angiographic correlates. *Ann Acad Med Singapore* 1996;25(2):196-9. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Yeon SB, Sabir A, Clouse M., et al. Delayed-enhancement cardiovascular magnetic resonance coronary artery wall imaging: comparison with multislice computed tomography and quantitative coronary angiography. *J Am Coll Cardiol* 2007;50(5):441-7. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Yildirim A, Kaynar G, Gulmez O., et al. Is it possible to increase the diagnostic value of exercise electrocardiography by post-exercise B-type natriuretic peptide levels? *Acta Cardiol* 2007;62(1):39-45. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Yilmaz MH, Yasar D, Albayram S,, et al. Coronary calcium scoring with MDCT: the radiation dose to the breast and the effectiveness of bismuth breast shield. *Eur J Radiol* 2007;61(1):139-43. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization.*

Yip GW, Chandrasekaran K, Miller TD,, et al. Feasibility of continuous venous infusion of SonoVue for qualitative assessment of reversible coronary perfusion defects in stress myocardial contrast echocardiography. *Int J Cardiovasc Imaging* 2003;19(6):473-81. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Yorgun H, Kaya EB, Hazirolan T,, et al. Prevalence of incidental pulmonary findings and early follow-up results in patients undergoing dual-source 64-slice computed tomography coronary angiography. *J Comput Assist Tomogr* 2010;34(2):296-301. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Yosefy C, Cantor A, Reisin L,, et al. The diagnostic value of QRS changes for prediction of coronary artery disease during exercise testing in women: false-positive rates. *Coron Artery Dis* 2004;15(3):147-54. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization.*

Yoshinaga K, Chow BJ, Williams K,, et al. What is the prognostic value of myocardial perfusion imaging using rubidium-82 positron emission tomography? *J Am Coll Cardiol* 2006;48(5):1029-39. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Yoshitani H, Takeuchi M, Mor-Avi V,, et al. Comparative diagnostic accuracy of multiplane and multislice three-dimensional dobutamine stress echocardiography in the diagnosis of coronary artery disease. *J Am Soc Echocardiogr* 2009;22(5):437-42. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Yoshitani H, Takeuchi M, Sakamoto K,, et al. Effect of one or more co-morbid conditions on diagnostic accuracy of coronary flow velocity reserve for detecting significant left anterior descending coronary stenosis. *Heart* 2005;91(10):1294-8. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Younis LT, Byers S, Shaw L,, et al. Prognostic importance of silent myocardial ischemia detected by intravenous dipyridamole thallium myocardial imaging in asymptomatic patients with coronary artery disease. *J Am Coll Cardiol* 1989;14(7):1635-41. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Yuda S, Khoury V, Marwick TH. Influence of wall stress and left ventricular geometry on the accuracy of dobutamine stress echocardiography. *J Am Coll Cardiol* 2002;40(7):1311-9. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Zafir N, Mats I, Solodky A,, et al. Prognostic value of stress myocardial perfusion imaging in octogenarian population. *J Nucl Cardiol* 2005;12(6):671-5. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Zafir N, Mats I, Solodky A,, et al. Myocardial perfusion profile in a young population with and without known coronary artery disease: comparison by gender. *Clin Cardiol* 2010;33(2):E39-43. *Full-text exclusion reason(s): Study does not compare one NIT to another or to diagnostic cardiac catheterization.*

Zaid G, Tanchilevitch A, Rivlin E,, et al. Diagnostic accuracy of serum B-type natriuretic peptide for myocardial ischemia detection during exercise testing with spect perfusion imaging. *Int J Cardiol* 2007;117(2):157-64. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup; No outcomes of interest.*

Zaid G, Yehudai D, Rosenschein U,, et al. Coronary artery disease in an asymptomatic population undergoing a multidetector computed tomography (mdct) coronary angiography. *Open Cardiovascular Medicine Journal* 2010;4:7-13. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Study does not compare one NIT to another or to diagnostic cardiac catheterization.*

Zakavi SR, Taherpour M, Saleh F,, et al. Electrocardiographic changes after dipyridamole infusion in patients undergoing myocardial perfusion imaging. *Nucl Med Commun* 2010;31(6):502-505. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Zaman MM, Haque SS, Siddique MA,, et al. Correlation between Severity of Coronary Artery Stenosis and Perfusion Defect Assessed by SPECT Myocardial Perfusion Imaging. *Mymensingh Med J* 2010;19(4):608-13. *Full-text exclusion reason(s): Full-text unobtainable.*

Zanco P, Zampiero A, Favero A,, et al. Myocardial technetium-99m sestamibi single-photon emission tomography as a prognostic tool in coronary artery disease: multivariate analysis in a long-term prospective study. *Eur J Nucl Med* 1995;22(9):1023-8. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Zellweger MJ, Dubois EA, Lai S,, et al. Risk stratification in patients with remote prior myocardial infarction using rest-stress myocardial perfusion SPECT: prognostic value and impact on referral to early catheterization. *J Nucl Cardiol* 2002;9(1):23-32. *Full-text exclusion reason(s): All women in the study are known to have CAD; Study does not compare one NIT to another or to diagnostic cardiac catheterization; Data for women not reported as a subgroup.*

Zerahn B, Jensen BV, Nielsen KD,, et al. Increased prognostic value of combined myocardial perfusion imaging and exercise electrocardiography in patients with coronary artery disease. *J Nucl Cardiol* 2000;7(6):616-22. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; All women in the study are known to have CAD; Data for women not reported as a subgroup.*

Zhang JJ, Liu T, Feng Y,, et al. Diagnostic value of 64-slice dual-source CT coronary angiography in patients with atrial fibrillation: Comparison with invasive coronary angiography. *Korean J Radiol* 2011;12(4):416-423. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Zhang LJ, Wu SY, Wang J,, et al. Diagnostic accuracy of dual-source CT coronary angiography: The effect of average heart rate, heart rate variability, and calcium score in a clinical perspective. *Acta Radiol* 2010;51(7):727-40. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Zhang ZH, Jin ZY, Li DJ,, et al. Non-invasive imaging of coronary artery with 16-slice spiral computed tomography. *Chin Med Sci J* 2004;19(3):174-9. *Full-text exclusion reason(s): No women with symptomatic chest pain, or results are not reported separately for symptomatic subgroup; Data for women not reported as a subgroup.*

Zhao Y, Wei JP, Hua Q,, et al. Comparison of multi-slice CT coronary artery imaging with coronary angiography. *Journal of Clinical Rehabilitative Tissue Engineering Research* 2008;12(44):8792-8796. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*

Zoghbi WA, Cheirif J, Kleiman NS,, et al. Diagnosis of ischemic heart disease with adenosine echocardiography. *J Am Coll Cardiol* 1991;18(5):1271-9. *Full-text exclusion reason(s): Data for women not reported as a subgroup.*