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STRESS ECHOCARDIOGRAPHY	
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### **GENERAL INFORMATION**

- It is an expectation that all patients receive care/services from a licensed clinician. All appropriate supporting documentation, including recent pertinent office visit notes, laboratory data, and results of any special testing must be provided. If applicable: All prior relevant imaging results and the reason that alternative imaging cannot be performed must be included in the documentation submitted.
- Where a specific clinical indication is not directly addressed in this guideline, medical necessity
  determination will be made based on widely accepted standard of care criteria. These criteria are
  supported by evidence-based or peer-reviewed sources such as medical literature, societal
  guidelines and state/national recommendations.

This guideline is for stress imaging, specifically Stress Echocardiography (SE) with appropriate preference for suitable alternatives, such as an exercise treadmill exam without imaging, when more suitable, unless otherwise stated (refer to <u>Overview section</u>).

INDICATIONS for STRES ECHO 1-3

SUSPECTED CORONARY ARTERY DISEASE (CAD)

Symptomatic patients without known CAD (use Diamond Forrester Table)

- Low or intermediate pretest probability, and electrocardiogram (ECG) is uninterpretable
- High pretest probability
- Repeat testing in patient with new or worsening symptoms and negative result at least one year ago AND meets one of the criteria above

### Asymptomatic patients without known CAD

- Previously unevaluated ECG evidence of possible myocardial ischemia including ischemic ST segment or T wave abnormalities (see Overview section)
- Previously unevaluated pathologic Q waves (<u>see Overview section</u>)
- Previously unevaluated complete left bundle branch block

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# ABNORMAL CALCIUM SCORES (CAC)1, 4-7

- ASYMPTOMATIC patient with a calcium score >400, not previously evaluated
- SYMPTOMATIC patient with prior CAC  $\geq$ 100

#### INCONCLUSIVE CAD EVALUATION AND OBSTRUCTIVE CAD REMAINS A CONCERN

- Exercise stress ECG with low-risk Duke treadmill score ≥5, but patient's current symptoms indicate an intermediate or high pretest probability
- Exercise stress ECG with an intermediate Duke treadmill score
- Intermediate coronary computed tomography angiography (CCTA) defined as:
  - 40 -70% lesion
- Coronary stenosis of unclear significance on previous coronary angiography<sup>1</sup>

### FOLLOW-UP OF PATIENTS POST CORONARY REVASCULARIZATION (PCI or CABG)8

 Asymptomatic, follow-up stress imaging (MPI or SE), at a minimum of 2 years post coronary artery bypass grafting (CABG), or percutaneous coronary intervention (PCI), whichever is later, is appropriate for patients with a history of silent ischemia or a history of a prior left main stent<sup>1</sup>

#### OR

- For patients with high occupational risk including any of the following:
  - Associated with public safety
  - Airline and boat pilots
  - Bus and train drivers
  - Bridge and tunnel workers/toll collectors
  - Police officers and firefighters
- New, recurrent, or worsening symptoms post coronary revascularization is an indication for stress imaging

### **FOLLOW-UP OF KNOWN CAD**

• Routine follow-up of asymptomatic or stable symptoms when last invasive or non-invasive assessment of coronary disease showed hemodynamically significant CAD (ischemia on stress test or FFR ≤ 0.80 or significant stenosis in a major vessel (≥ 50% left main coronary artery or ≥ 70% LAD, LCX, RCA)), over two years ago without intervening coronary revascularization, is an appropriate indication for stress imaging (MPI or SE)

### SPECIAL DIAGNOSTIC CONDITIONS REQUIRING CORONARY EVALUATION

 Prior acute coronary syndrome (with documentation in MD notes), within last three months, without a prior stress test or coronary angiography performed since that time



- Newly diagnosed systolic heart failure or diastolic heart failure, with reasonable suspicion of cardiac ischemia (prior events, risk factors), unless invasive coronary angiography is immediately planned<sup>4,8</sup>
- Ventricular arrhythmias:
  - Sustained ventricular tachycardia (VT) > 100 bpm, ventricular fibrillation (VF), or exercise-induced VT, when invasive coronary arteriography has not been performed<sup>9</sup>
  - Nonsustained VT, multiple episodes, each ≥ 3 beats at ≥ 100 bpm, frequent PVCs (defined as greater than or equal to 30/hour on remote monitoring), when an exercise ECG cannot be performed<sup>9</sup>
- For intermediate and high-risk global patients who require initiation of Class IC antiarrhythmic drugs. It can be performed annually thereafter until discontinuation of drug use<sup>10</sup>
- Hemodynamic assessment of ischemia in one of the following documented conditions:
  - Anomalous coronary arteries in an asymptomatic individual without prior stress echocardiography<sup>11</sup>;
  - Myocardial bridging of a coronary artery<sup>12</sup>
- Coronary aneurysms in Kawasaki's disease<sup>13</sup>
- Following radiation therapy to the anterior or left chest, at 5 years post initiation and every 5 years thereafter<sup>14</sup>

### **CHRONIC VALVULAR DISEASE**

### Evaluation with Inclusion of Doppler<sup>15-18</sup>

- For the evaluation of aortic stenosis and flow (contractile) reserve in symptomatic patients with severe aortic stenosis by calculated valve area, low flow / low gradient, and ejection fraction < 50%</li>
- For evaluation of asymptomatic moderate or severe aortic stenosis (AS) for measurement of changes in valve hemodynamics
- Non-severe aortic regurgitation (AR) with symptoms: Assessment of functional capacity and to assess for other causes of symptoms<sup>8, 19</sup>
- For evaluation of mitral stenosis (MS) if there is:
  - Exertional shortness of breath which suggests the amount of MS is worse than is seen on the resting echocardiogram
- For evaluation for mitral regurgitation (MR) if there is:
  - Exertional shortness of breath which suggests the amount of MR is worse than is seen on the resting echocardiogram; OR
  - The echocardiogram is not able to distinguish whether the MR is moderate or severe in a patient that is asymptomatic



- For symptomatic patients with HCM, who do not have resting or provocable outflow tract gradient ≥50 mm Hg on TTE, for detection and quantification of dynamic LVOT obstruction<sup>20</sup>
- For asymptomatic patients with HCM who do not have a resting or provocable outflow tract gradient ≥ 50 mm Hg on TTE (Class 2A)

#### DIASTOLIC FUNCTION

 For unexplained dyspnea and suspected heart failure with preserved LVEF<sup>19</sup> (HFpEF) with normal or equivocal diastolic function on resting images

# PRIOR TO ELECTIVE NON-CARDIAC SURGERY<sup>2, 21-23</sup>

- An intermediate or high risk surgery with of one or more risk factors (see below), AND documentation of an inability to walk (or <4 METs) AND there has not been an imaging stress test within 1 year<sup>21, 23, 24</sup>
  - Risk factors: history of ischemic heart disease, history of congestive heart failure, history of cerebrovascular disease, preoperative treatment with insulin, and preoperative serum creatinine >2.0 mg/dL.
  - Surgical Risks:
    - High risk surgery: Aortic and other major vascular surgery, peripheral vascular surgery, anticipated prolonged surgical procedures associated with large fluid shifts and/or blood loss
    - Intermediate risk surgery: Carotid endarterectomy, head and neck surgery, intraperitoneal and intrathoracic surgery, orthopedic surgery, prostate surgery
    - Low risk surgery: Endoscopic procedures, superficial procedure, cataract surgery, breast surgery
- Planning for any organ or stem cell transplantation is an indication for preoperative stress imaging, if there has not been a conclusive stress evaluation, CTA, or heart catheterization within the past year, at the discretion of the transplant service. <sup>2, 25</sup>

#### POST CARDIAC TRANSPLANTATION

- Annually, for the first five years post cardiac transplantation, in a patient not undergoing invasive coronary arteriography
- After the first five years post cardiac transplantation, patients with documented transplant coronary vasculopathy can be screened annually unless invasive coronary arteriography is planned

## **BACKGROUND**



Stress echocardiography (SE) refers to ultrasound imaging of the heart during exercise electrocardiography (ECG) testing, during which visualized wall motion abnormalities can provide evidence of potential significant coronary artery disease (CAD).

While drug-induced stress with dobutamine can be an alternative to exercise stress testing in patients who are unable to exercise, this guideline does not require use of this modality. Hence, reference in this document to SE predominantly refers to exercise stress echocardiography.

Although SE provides comparable accuracy without radiation risk, relative to myocardial perfusion imaging (MPI), scenarios which do not permit effective use of SE might be better suited for stress imaging with MPI, cardiovascular magnetic resonance imaging (CMR) or positron emission tomography (PET), or coronary computed tomography angiography (CCTA).

## Stable patients without known CAD fall into 2 categories<sup>1-3</sup>:

- Asymptomatic patients, for whom Global Risk of CAD events can be determined from coronary risk factors using calculators available online (see Websites for <u>Global</u> <u>Cardiovascular Risk Calculators</u> section)
- **Symptomatic patients,** for whom we estimate the Pretest Probability that their chest-related symptoms are due to clinically significant CAD (see below):

### The 3 Types of Chest Pain or Discomfort:

- Typical Angina (Definite) is defined as including all 3 of these characteristics:
  - Substernal chest pain or discomfort with characteristic quality and duration, such as:
    - Pressure-like
    - Radiating
    - Dull or aching
  - Provoked by exertion or emotional stress
  - Relieved by rest and/or nitroglycerine
- Atypical Angina (Probable) has only 2 of the above characteristics
- Nonanginal Chest Pain/Discomfort has only 0-1 of the above characteristics

The medical record should provide enough detail to establish the type of chest pain. From those details, the Pretest Probability of obstructive CAD is estimated from the <u>Diamond</u> <u>Forrester Table</u> below, recognizing that in some cases multiple additional coronary risk factors could increase pretest probability<sup>1, 3</sup>:



#### **Diamond Forrester Table**

Age (Years)	Gender	Typical/Definite Angina Pectoris	Atypical/Probable Angina Pectoris	Nonanginal Chest Pain
≤ 39	Men	Intermediate	Intermediate	Low
≥ 39	Women	Intermediate	Very low	Very low
40 – 49	Men	High	Intermediate	Intermediate
40 – 49	Women	Intermediate	Low	Very low
50 – 59	Men	High	Intermediate	Intermediate
30 – 39	Women	Intermediate	Intermediate	Low
≥ 60	Men	High	Intermediate	Intermediate
2 00	Women	High	Intermediate	Intermediate

- Very low: < 5% pretest probability of CAD, usually not requiring stress evaluation<sup>3</sup>
- Low: 5 10% pretest probability of CAD
- Intermediate: 10% 90% pretest probability of CAD
- High: > 90% pretest probability of CAD

### **OVERVIEW**

MPI may be performed without diversion to SE in any of the following<sup>1, 26</sup>:

- Inability to exercise
  - Physical limitations precluding ability to exercise for at least 3 full minutes of Bruce protocol
  - Limited functional capacity (< 4 metabolic equivalents) such as one of the following:
    - Cannot take care of their activities of daily living (ADLs) or ambulate
    - Cannot walk 2 blocks on level ground
    - Cannot climb 1 flight of stairs
    - Cannot vacuum, dust, do dishes, sweep, or carry a small grocery bag
- Other Comorbidities
  - Severe chronic obstructive pulmonary disease with pulmonary function test (PFT) documentation, severe shortness of breath on minimal exertion, or requirement of home oxygen during the day
  - Poorly controlled hypertension, with systolic BP > 180 or Diastolic BP > 120 (and clinical urgency not to delay MPI)
- ECG and Echo-Related Baseline Findings
  - Prior cardiac surgery (coronary artery bypass graft or valvular)
  - Documented poor acoustic imaging window
  - Left ventricular ejection fraction ≤ 40%
  - Pacemaker or ICD
  - Persistent atrial fibrillation
  - Resting wall motion abnormalities that would make SE interpretation difficult



- Complete LBBB
- Risk-related scenarios
  - High pretest probability in suspected CAD
  - Intermediate or high global risk in patients requiring type IC antiarrhythmic drugs (prior to initiation of therapy and annually)
  - Arrhythmia risk with exercise
- Previously unevaluated pathologic Q waves (in two contiguous leads) defined as the following:
  - o > 40 ms (1 mm) wide
  - o > 2 mm deep
  - > 25% of depth of QRS complex

### **ECG Stress Test Alone versus Stress Testing with Imaging**

Prominent scenarios suitable for an ECG stress test **WITHOUT** imaging (i.e., exercise treadmill ECG test) are inferred from the guidelines presented above, often requiring that the patient can exercise for at least 3 minutes of Bruce protocol with achievement of near maximal heart rate **AND** has an interpretable ECG for ischemia during exercise<sup>1</sup>:

- The (symptomatic) low or intermediate pretest probability patient who is able to exercise and has an interpretable ECG
- The patient who is under evaluation for exercise-induced arrhythmia<sup>9</sup>
- For the evaluation of syncope or presyncope during exertion<sup>27</sup>
- The patient who requires an entrance stress test ECG for a cardiac rehab program or for an exercise prescription.

# **Duke Exercise ECG Treadmill Score<sup>28</sup>**

Calculates risk from ECG treadmill alone:

- The equation for calculating the Duke treadmill score (DTS) is: DTS = exercise time in minutes (5 x ST deviation in mm or 0.1 mV increments) (4 x exercise angina score), with angina score being 0 = none, 1 = non-limiting, and 2 = exercise-limiting.
- The score typically ranges from 25 to + 15. These values correspond to low-risk (with a score of ≥ + 5), intermediate risk (with scores ranging from 10 to + 4), and high-risk (with a score of ≤ -11) categories.

An uninterpretable baseline ECG includes<sup>3</sup>:

- ST segment depression 1 mm or more; (not for non-specific ST- T wave changes)
- Ischemic looking T wave -- at least 2.5 mm inversions (excluding V1 and V2)
- LVH, pre-excitation pattern such as WPW, a ventricular paced rhythm, or left bundle branch block
- Digitalis use



 Resting HR under 50 bpm on a medication, such as beta-blockers or calcium channel blockers, that is required for patient's treatment and cannot be stopped, with an anticipated suboptimal workload

### **Global Risk of Cardiovascular Disease**

**Global risk** of CAD is defined as the probability of manifesting cardiovascular disease over the next 10 years and refers to **asymptomatic** patients without known cardiovascular disease. It should be determined using one of the risk calculators below. A high risk is considered greater than a 20% risk of a cardiovascular event over the ensuing 10 years. High global risk by itself generally lacks scientific support as an indication for stress imaging. There are rare exemptions, such as patients requiring IC antiarrhythmic drugs, who might require coronary risk stratification prior to initiation of the drug.

- CAD Risk—Low
  - 10-year absolute coronary or cardiovascular risk less than 10%.
- CAD Risk—Moderate
  - 10-year absolute coronary or cardiovascular risk between 10% and 20%.
- CAD Risk—High
  - 10-year absolute coronary or cardiovascular risk of greater than 20%.

### Websites for Global Cardiovascular Risk Calculators\*

	iovasculai Nisk Calculators
Risk Calculator	Link to Online Calculator
Framingham	https://reference.medscape.com/calculator/framingham-
Cardiovascular Risk	<u>cardiovascular-disease-risk</u>
Reynolds Risk Score	http://www.reynoldsriskscore.org/
Can use if no diabetes	
Unique for use of	
family history	
Pooled Cohort	http://clincalc.com/Cardiology/ASCVD/PooledCohort.aspx?example
Equation	
ACC/AHA Risk	http://tools.acc.org/ASCVD-Risk-Estimator/
Calculator	
MESA Risk Calculator	https://www.mesa-
With addition of	nhlbi.org/MESACHDRisk/MesaRiskScore/RiskScore.aspx
Coronary Artery	
Calcium Score, for	
CAD-only risk	

<sup>\*</sup>Patients who have known CAD are already at high global risk and are not applicable to the calculators.<sup>29-33</sup>

# Definitions of Coronary Artery Disease<sup>2, 3, 5, 34, 35</sup>



- Percentage stenosis refers to the reduction in diameter stenosis when angiography is the method and refers to cross-sectional narrowing when IVUS (intravascular ultrasound) is the method of determination
- Coronary artery calcification is a marker of risk, as measured by Agatston score on coronary artery calcium imaging. Its incorporation into Global Risk can be achieved by using the MESA risk calculator.
- Ischemia-producing disease (also called hemodynamically or functionally significant disease, for which revascularization might be appropriate), generally implies at least one of the following:
  - O Suggested by percentage diameter stenosis  $\geq$  70% by angiography; intermediate lesions are  $50 69\%^{36}$
  - For a left main artery, suggested by a percentage stenosis  $\geq$  50% or minimum lumen cross-sectional area on IVUS  $\leq$  6 square mm<sup>3, 35, 37</sup>
  - FFR (fractional flow reserve)  $\leq$  0.80 for a major vessel<sup>35, 37</sup>
- FFR (fractional flow reserve) is the distal to proximal pressure ratio across a coronary lesion during maximal hyperemia induced by either intravenous or intracoronary adenosine. Less than or equal to 0.80 is considered a significant reduction in coronary flow

# Anginal Equivalent<sup>3, 27, 38</sup>

Development of an anginal equivalent (e.g., shortness of breath, fatigue, or weakness) either with or without prior coronary revascularization should be based upon the documentation of reasons to suspect that symptoms other than chest discomfort are not due to other organ systems (e.g., dyspnea due to lung disease, fatigue due to anemia). This may include respiratory rate, oximetry, lung exam, etc. (as well as d-dimer, chest CT(A), and/or PFTs, when appropriate), and then incorporated into the evaluation of coronary artery disease as would chest discomfort. Syncope per se is not an anginal equivalent.



#### **Abbreviations**

AAD Antiarrhythmic drug
ADLs Activities of daily living

BSA Body surface area in square meters
CABG Coronary artery bypass grafting surgery

CAC Coronary artery calcium CAD Coronary artery disease

CCTA Coronary computed tomography angiography
CMR Cardiovascular magnetic resonance imaging
CT(A) Computed tomography (angiography)

DTS Duke Treadmill Score
ECG Electrocardiogram
FFR Fractional flow reserve

HCM Hypertrophic cardiomyopathy
IVUS Intravascular ultrasound
LBBB Left bundle-branch block

LVEF Left ventricular ejection fraction
LVH Left ventricular hypertrophy
LVOT Left ventricular outflow tract

MESA Multi-Ethnic Study of Atherosclerosis

MET Estimated metabolic equivalent of exercise

MI Myocardial infarction

MPI Myocardial perfusion imaging

MR Mitral regurgitation MS Mitral stenosis

PCI Percutaneous coronary intervention

PET Positron emission tomography

PFT Pulmonary function test

PVCs Premature ventricular contractions

SE Stress echocardiography

TTE Transthoracic echocardiography

VT Ventricular tachycardia VF Ventricular fibrillation WPW Wolff-Parkinson-White



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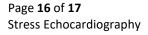


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# **POLICY HISTORY**

Date	Summary
May 2023	<ul> <li>Removed time limitation "within past two years" for further evaluation inconclusive prior CAD evaluation</li> <li>Added coronary stenosis of unclear significance on coronary angiography</li> <li>Added evaluation of asymptomatic moderate or severe aortic stenosis (AS) and aortic regurgitation (AR) for measurement of changes in valve hemodynamics</li> <li>Added evaluation symptomatic patients with suspected diastolic dysfunction</li> <li>Added statement on clinical indications not addressed in this guideline</li> </ul>
February 2022	<ul> <li>Moved the sentence regarding utilization of suitable alternatives such as Stress Echocardiography and MPI to the General Information section</li> <li>Clarified "intermediate lesions are 50-69%" for ischemia-producing disease</li> <li>Placed Link to Overview Section in General Information</li> <li>Deleted the requirement for diabetes when calcium score &gt; 400 for stress imaging</li> <li>Added Calcium score section:</li> <li>Added stress imaging approval for calcium score &gt; 100 with symptoms consistent with low to intermediate pretest probability</li> <li>Changed preoperative guideline to include intermediate risk surgery with one or more risk factors AND documentation of an inability to walk (or &lt;4 METs) AND there has not been an imaging stress test within 1 year</li> <li>Changed solid organ transplant guideline to include stem cell transplant and "any" organ transplant</li> <li>Added definition of surgical risk to preop guidelines</li> <li>In Background section clarified the requirement for description of chest pain by adding sentence "The medical record should provide enough detail to establish the type of chest pain. "</li> <li>Added definition of Q waves</li> <li>Deleted sentence regarding calcium scoring within the Global Risk Section</li> <li>Deleted sentence regarding using calcium score solely for risk stratification</li> <li>Deleted IFR references</li> </ul>





### Reviewed / Approved by NIA Clinical Guideline Committee

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