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# Perioperative Cardiovascular Risk Assessment and Management for Noncardiac Surgery
## A Review

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- ⊞ Author Audio Interview
- ⊞ Supplemental content
- ⊞ CME Quiz at jamacmelookup.com

**IMPORTANCE** Perioperative cardiovascular complications occur in 3% of hospitalizations for noncardiac surgery in the US. This review summarizes evidence regarding cardiovascular risk assessment prior to noncardiac surgery.

**OBSERVATIONS** Preoperative cardiovascular risk assessment requires a focused history and physical examination to identify signs and symptoms of ischemic heart disease, heart failure, and severe valvular disease. Risk calculators, such as the Revised Cardiac Risk Index, identify individuals with low risk (<1%) and higher risk (≥1%) for perioperative major adverse cardiovascular events during the surgical hospital admission or within 30 days of surgery. Cardiovascular testing is rarely indicated in patients at low risk for major adverse cardiovascular events. Stress testing may be considered in patients at higher risk (determined by the inability to climb ≥2 flights of stairs, which is <4 metabolic equivalent tasks) if the results from the testing would change the perioperative medical, anesthesia, or surgical approaches. Routine coronary revascularization does not reduce perioperative risk and should not be performed without specific indications independent of planned surgery. Routine perioperative use of low-dose aspirin (100 mg/d) does not decrease cardiovascular events but does increase surgical bleeding. Statins are associated with fewer postoperative cardiovascular complications and lower mortality (1.8% vs 2.3% without statin use; $P < .001$) in observational studies, and should be considered preoperatively in patients with atherosclerotic cardiovascular disease undergoing vascular surgery. High-dose β-blockers (eg, 100 mg of metoprolol succinate) administered 2 to 4 hours prior to surgery are associated with a higher risk of stroke (1.0% vs 0.5% without β-blocker use; $P = .005$) and mortality (3.1% vs 2.3% without β-blocker use; $P = .03$) and should not be routinely used. There is a greater risk of perioperative myocardial infarction and major adverse cardiovascular events in adults aged 75 years or older (9.5% vs 4.8% for younger adults; $P < .001$) and in patients with coronary stents (8.9% vs 1.5% for those without stents; $P < .001$) and these patients warrant careful preoperative consideration.

**CONCLUSIONS AND RELEVANCE** Comprehensive history, physical examination, and assessment of functional capacity during daily life should be performed prior to noncardiac surgery to assess cardiovascular risk. Cardiovascular testing is rarely indicated in patients with a low risk of major adverse cardiovascular events, but may be useful in patients with poor functional capacity (<4 metabolic equivalent tasks) undergoing high-risk surgery if test results would change therapy independent of the planned surgery. Perioperative medical therapy should be prescribed based on patient-specific risk.

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Approximately 17.2 million surgeries are performed annually in the US.[1] Multiple cardiovascular risk factors such as hypertension and hyperlipidemia are present in 45% of patients aged 45 years or older undergoing noncardiac surgery, and nearly 25% have a history of atherosclerotic cardiovascular disease.[2] The incidence of perioperative cardiovascular events is related to the risk for cardiovascular events in the individual patient before surgery. In a retrospective study[3] of more than 10 million hospitalizations for noncardiac surgery in adults across the US, the combined rate of perioperative death, myocardial infarction, and ischemic stroke was 3.0%. Myocardial injury, defined as an elevated troponin level above the 99th percentile, occurs in up to 20% of patients after noncardiac surgery.[4,5] This review summarizes evidence regarding risk assessment, testing, and optimal medical therapy to reduce perioperative cardiovascular risk prior to noncardiac surgery (**Box 1**).

## Methods

We searched the MEDLINE database (using PubMed) and the Cochrane Library for English-language publications from January 1, 1949, through January 27, 2020, related to the evaluation of perioperative cardiovascular risk prior to noncardiac surgery. Clinical practice guidelines, randomized clinical trials, and meta-analyses of observational studies and trials were prioritized for review. Relevant references cited by identified articles were included. Included publications were mutually agreed upon by the authors and selected for clinical importance with consideration of the potential relevance for a general medical readership (eMethods in the Supplement).

## Estimating Perioperative Risk

Evaluating perioperative risk begins with a focused history and cardiovascular physical examination. The history should identify cardiovascular conditions associated with perioperative major adverse cardiovascular events (MACE), including history of ischemic heart disease,[6] coronary stent,[7] heart failure,[8-10] arrhythmias,[10] valvular heart disease,[11] systemic hypertension,[12] and pulmonary hypertension.[13,14] Cardiovascular disease risk factors, such as chronic kidney disease and diabetes, are associated with up to a 3-fold increased risk of cardiac events.[6]

Physicians should ask patients whether they can perform workloads of 4 or greater metabolic equivalent tasks (METs) without symptomatic limitation (eTable 1 in the Supplement), consisting of walking up a hill or climbing up 2 or more flights of stairs. Inability

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**Box 1. Questions Commonly Asked When Evaluating Perioperative Risk**

**1. Which risk scores provide the best discrimination of perioperative risk?**

The 6-component Revised Cardiac Risk Index is relatively simple to use. One point is assigned for each of the following: ischemic heart disease, cerebrovascular disease, heart failure, insulin-dependent diabetes, chronic kidney disease (serum creatinine level ≥2.0 mg/dL), and high-risk surgery (intraperitoneal, intrathoracic, or vascular). The 21-component National Surgical Quality Improvement Program universal surgical risk calculator is more complex but may provide better predictive discrimination.

**2. Should preoperative stress testing be routinely performed prior to noncardiac surgery?**

Routine cardiac stress testing is not indicated for low-risk patients or for high-risk patients who are able to walk up a hill or climb up 2 or more flights of stairs without difficulty. Testing may be considered for patients with unknown or poor functional capacity who may have high cardiovascular risk. Despite the established risks of coronary artery disease in surgical patients, coronary revascularization prior to surgery did not improve perioperative outcomes in a randomized trial. Thus, stress testing should only be considered if the results would change perioperative medical, anesthesia, or surgical approaches.

**3. Can β-blockers and statins prevent cardiovascular complications of noncardiac surgery?**

Perioperative use of β-blockers confers some theoretical advantages in reducing mismatch in myocardial oxygen supply and demand. However, high-dose extended-release metoprolol succinate (100 mg/d) initiated immediately prior to surgery is associated with increased perioperative stroke and mortality in randomized trials. Statin therapy administered during hospitalization for surgery is associated with reduced cardiovascular risk in observational data sets. However, randomized trials with 80 mg/d of atorvastatin vs placebo taken within 18 hours before surgery did not clearly demonstrate benefit. Statins should be considered preoperatively in patients with atherosclerotic cardiovascular disease and may be considered in patients with clinical risk factors undergoing higher-risk surgery.

**4. Should antithrombotic and anticoagulation therapy be discontinued prior to surgery?**

Routine administration of perioperative antiplatelet therapy prior to noncardiac surgery is not recommended because it is not associated with benefit and results in an increased risk of bleeding. Low-dose aspirin may be appropriate for a subset of patients when ischemic risks outweigh the bleeding risks, such as for patients with coronary artery stents. For patients taking warfarin or a direct oral anticoagulant for stroke prevention in atrial fibrillation, perioperative interruption of oral anticoagulation is safe, and bridging with heparin should not be routinely performed. Patients with mechanical mitral valves and those at increased risk for thrombotic events with mechanical aortic valves should receive bridging anticoagulation with heparin prior to noncardiac surgery.

**5. How soon after coronary stent implantation may a patient safely undergo noncardiac surgery?**

Individuals who require surgery within 1 year after percutaneous coronary intervention are at increased risk of perioperative events compared with those without coronary stents. Ischemic risks are inversely related to the time interval between stent placement and noncardiac surgery. Patients who undergo coronary stent placement should have surgery delayed until the risks associated with delaying surgery outweigh the risks of thrombosis that are associated with cessation of dual antiplatelet therapy. Elective noncardiac surgery should be delayed for at least 30 days after bare metal stent implantation and 12 months after drug-eluting stent placement, although more recent evidence suggests that a delay of 3 to 6 months may be safe.

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to do so for any reason is independently associated with a 2-fold increased risk of perioperative complications.[15] Exertional chest pain, dyspnea, orthopnea, palpitations, recent syncope, and physical examination findings, such as murmurs (any diastolic or grade ≥3/6 systolic), gallops, jugular venous distention, or edema, may indicate cardiovascular disease. Ongoing, high-risk cardiac conditions such as acute coronary syndromes or decompensated heart failure are generally contraindications to noncardiac surgery and require additional evaluation (**Box 2**).

Type of surgery is also associated with the degree of risk for MACE (**Box 3**). By expert consensus, noncardiac surgeries with less than a 1% risk for MACE, such as cataract surgery and many types of cosmetic or plastic surgery, are considered low risk.[16] Vascular (7.7%), thoracic (6.5%), transplant (6.2%), and general (3.9%) surgeries are associated with the highest incidence of MACE.[3,6] Use of minimally invasive, laparoscopic, and endovascular techniques may attenuate cardiovascular risk.[17,18] In a randomized trial of open vs endovascular surgical abdominal aortic aneurysm repair, 30-day operative mortality was 4.3% in participants assigned to conventional open surgery vs 1.8% in those assigned to endovascular treatment.[19]

Classification systems and risk scores can help estimate perioperative risk.[6,20-24] The American Society of Anesthesiologists (ASA) Physical Status Classification System, for example, classifies patients into categories according to their overall health status and is independently associated with surgical outcomes. In a prospective study of 6301 patients, healthy patients (ASA class I) had a 0.1% risk of cardiac complications and mortality, whereas patients with "severe systemic disease that is a constant threat to life" (ASA class IV) had an 18% risk.[25] Cardiovascular risk scores commonly used include the Revised Cardiac Risk Index[6,26] and the National Surgical Quality Improvement Program perioperative myocardial infarction and cardiac arrest risk calculator and the universal surgical risk calculator (**Table**).[21,22] These scores provide estimates of cardiovascular risk based on perioperative factors. For example, to calculate the Revised Cardiac Risk Index (range, 0-6; 6 = worst), 1 point is assigned for each of the following: ischemic heart disease, cerebrovascular disease, heart failure, insulin-dependent diabetes, chronic kidney disease (serum creatinine level ≥2.0 mg/dL), and high-risk surgery (intraperitoneal, intrathoracic, or vascular). Patients with a Revised Cardiac Risk Index of 0 have an approximate risk of 0.4% for major cardiovascular complications, whereas those with an index of 3 or greater have an approximate risk of 10%. In a pooled analysis of 24 validation studies, the Revised Cardiac Risk Index had modest risk discrimination for cardiac events in patients undergoing noncardiac surgery (receiver operating characteristic curve, 0.75) and had poorer discrimination in patients undergoing vascular surgery (receiver operating characteristic curve, 0.64).[26] The 21-component National Surgical Quality Improvement Program universal surgical risk calculator may provide superior predictive discrimination.

## Preoperative Cardiovascular Testing
An algorithm for perioperative cardiovascular risk stratification appears in **Figure 1**,[16,18] but has not been tested in a randomized clinical trial. Perioperative guideline recommendations from the American Heart Association and the American College of Cardiology (AHA/ACC), the Canadian Cardiovascular Society, and the European Society of Cardiology appear in eTable 2 in the Supplement.

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**Box 2. High-Risk Cardiac Conditions Considered Contraindications to Noncardiac Surgery**

**Contraindications to Noncardiac Surgery**
- Acute coronary syndrome
- Acute decompensated heart failure
- Tachyarrhythmias or bradyarrhythmias associated with hypotension or requiring urgent medical attention (eg, ventricular tachycardia or high-grade atrioventricular block)
- Symptomatic, severe aortic stenosis (mean gradient >40 mm Hg or peak velocity >4 m/s)

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**Box 3. Cardiovascular Risk Classification and Examples of Surgery Types[3,6]**

**Level of Risk for Major Adverse Cardiovascular Events or Death**

**<1% Risk**
Cataract surgery

Cosmetic or plastic surgery

**≥1% Risk**
Orthopedic surgery

Otolaryngology surgery

Genitourinary surgery

**≥3% Risk**
General abdominal or intraperitoneal surgery

Neurosurgery

**≥5% Risk**
Suprainguinal and peripheral vascular surgery

Thoracic surgery

Transplant surgery

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### 12-Lead Electrocardiographic Testing
Preoperative 12-lead electrocardiographic (ECG) testing defines the cardiac rhythm, identifies clinically silent cardiovascular disease such as prior Q-wave myocardial infarction, and provides a baseline for postoperative comparison. Among patients with coronary artery disease (CAD) undergoing major surgery, preoperative ST-segment depressions greater than 0.5 mm are associated with increased risk of postoperative death or myocardial infarction (event rate of 11.2% in patients with ST-segment depressions vs 2.6% in those without such depressions; P = .001).[27,28] However, an ECG provides little benefit prior to low-risk surgery such as cataract surgery and cosmetic or plastic surgery.[29-31] Therefore, preoperative 12-lead ECG is reasonable in patients with a history of CAD, arrhythmias, peripheral artery disease, cerebrovascular disease, or structural heart disease scheduled for higher-risk surgery.[16] A 12-lead ECG also is reasonable prior to higher-risk surgeries such as major abdominal or thoracic procedures even among those without symptoms of cardiovascular disease.[16]

### Transthoracic Echocardiography
Echocardiography is a noninvasive imaging modality that evaluates left ventricular function and valvular heart disease. In an observational study of 570 patients undergoing noncardiac surgery, elevated aortic valve gradients of 40 mm Hg or greater (odds ratio

**Table. Risk Scores and Calculators**

| | Goldman Index of Cardiac Risk,[20] 1977 | Revised Cardiac Risk Index,[6] 1999 | National Surgical Quality Improvement Program | | Geriatric-Sensitive Perioperative Cardiac Risk Index,[24] 2017 | Cardiovascular Risk Index,[23] 2019 |
| | | | Risk calculators | | | |
| | | | Perioperative MI and cardiac arrest,[21] 2011 | Universal surgical,[22] 2013 | | |
|---|---|---|---|---|---|---|
| Criteria | • Aged >70 y (5 points)<br>• Had an MI within 6 mo (10 points)<br>• Jugular venous distention or a third heart sound on auscultation (11 points)<br>• ≥5 PVCs/min (7 points)<br>• Nonsinus rhythm or PAC on preoperative ECG (7 points)<br>• Aortic stenosis (3 points)<br>• Intraperitoneal, intrathoracic, or aortic surgery (3 points)<br>• Any emergency surgery (4 points) | • Ischemic heart disease (1 point)<br>• Cerebrovascular disease (1 point)<br>• History of congestive heart failure (1 point)<br>• Insulin therapy for diabetes (1 point)<br>• Serum creatinine level ≥2.0 mg/dL (1 point)<br>• Planned high-risk procedure (intraperitoneal, intrathoracic, or vascular surgery) (1 point) | • Age<br>• ASA class<br>• Preoperative function<br>• Creatinine level<br>• Procedure type: anorectal; aortic; bariatric; brain; breast; cardiac; ear, nose, or throat; foregut or hepatopancreatobiliary; gallbladder, appendix, adrenal, or spleen; intestinal; neck; obstetric or gynecologic; orthopedic; other abdomen; peripheral vascular; skin; spine; thoracic; urological; or vein | • Age group<br>• Sex<br>• ASA class<br>• Functional status<br>• Emergency case<br>• Steroid use for chronic condition<br>• Ascites within 30 d preoperatively<br>• System sepsis within 48 h preoperatively<br>• Required ventilator<br>• Disseminated cancer<br>• Diabetes<br>• Hypertension requiring medication<br>• Prior cardiac event<br>• Congestive heart failure within 30 d preoperatively<br>• Dyspnea<br>• Current smoker within 1 y<br>• History of COPD<br>• Dialysis<br>• Acute kidney failure<br>• BMI<br>• CPT-specific linear risk | • Age<br>• Sex<br>• ASA class<br>• High-risk surgery<br>• History of heart failure<br>• Stroke<br>• Required insulin<br>• Diabetes<br>• Dialysis<br>• Medications for hypertension<br>• Current tobacco use<br>• History of COPD<br>• Functional status (partially vs totally dependent)<br>• Creatinine level<br>• Surgical category<br>• Dyspnea<br>• BUN level<br>• Laparoscopic surgery | • Age ≥75 y (1 point)<br>• History of heart disease (1 point)<br>• Symptoms of angina or dyspnea (1 point)<br>• Hemoglobin level <12 mg/dL (1 point)<br>• Vascular surgery (1 point)<br>• Emergency surgery (1 point) |
| Score range | • Class I: 0-5 points (lowest risk)<br>• Class II: 6-12 points<br>• Class III: 13-25 points<br>• Class IV: ≥26 points (highest risk) | • Class I: 0 points (lowest risk)<br>• Class II: 1 point<br>• Class III: 2 points<br>• Class IV: ≥3 points (highest risk) | 0%-100% (0%, lowest risk; 100%, highest risk) | 0%-100% (0%, lowest risk; 100%, highest risk) | 0%-100% (0%, lowest risk; 100%, highest risk) | • 0 points (lowest risk)<br>• 1 point<br>• 2 points<br>• 3 points<br>• >3 points (highest risk) |
| Threshold denoting elevated risk | ≥Class II (≥6 points) | >1 point | >1% | >1% | >1% | ≥2 points |
| Outcome | Intraoperative or postoperative MI, pulmonary edema, VT, cardiac death | MI, pulmonary edema, ventricular fibrillation, complete heart block, cardiac death | Intraoperative or postoperative MI or cardiac arrest within 30 d | Cardiac arrest, MI, all-cause mortality within 30 d | Cardiac arrest, MI, all-cause mortality within 30 d | Death, MI, or stroke at 30 d |
| Derivation population | 1001 | 1422 | 211 410 | 1 414 006 | 584 931 | 3284 |
| Set ROC | | | | | | |
| Derivation | 0.61 | 0.76 | 0.88 | 0.90 (Cardiac arrest or MI); 0.94 (mortality) | 0.90 | |
| Validation | 0.70 | 0.81; 0.75[a] | 0.87[b] | 0.88 (Cardiac arrest or MI); 0.94 (mortality)[b] | 0.83 (0.76 in adults aged ≥65 y)[b] | 0.82[b] |

Abbreviations: ASA, American Society of Anesthesiologists; BMI, body mass index; BUN, blood (serum) urea nitrogen; COPD, chronic obstructive pulmonary disease; CPT, *Current Procedural Terminology*; ECG, electrocardiogram; MI, myocardial infarction; PAC, premature atrial contraction; PVC, premature ventricular contraction; ROC, receiver operating characteristic curve; VT, ventricular tachycardia.

a Pooled validation studies assessing the performance in mixed noncardiac surgery.[26]

b Validated using the National Surgical Quality Improvement Program database. The risk calculators are available at https://riskcalculator.facs.org/RiskCalculator/PatientInfo.jsp.

Figure 1. A Proposed Algorithm for Perioperative Cardiovascular Risk Assessment
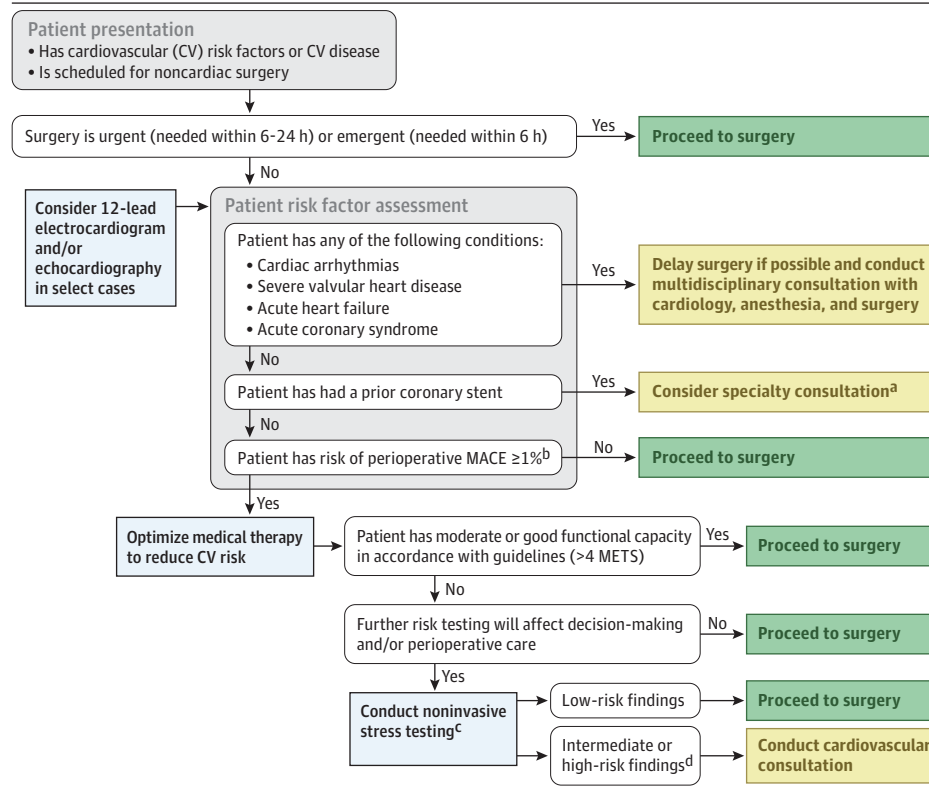


The algorithm has not been validated. MACE indicates major adverse cardiovascular events; METs metabolic equivalent tasks.

[a] Perioperative considerations during consulation shown in Figure 2.

[b] Risk of perioperative MACE as determined by a clinical risk calculator.

[c] Testing options include: (1) exercise electrocardiographic stress testing without myocardial imaging; or (2) stress testing (exercise or pharmacological) with imaging such as echocardiography, nuclear perfusion via single-photon emission computed tomography, positron emission tomography, or cardiac magnetic resonance imaging.

[d] Intermediate or high-risk findings by stress testing may include moderate to severe myocardial ischemia, ischemia provoked at a low workload, a hypotensive response to exercise, transient ischemic dilatation, and ventricular arrhythmias during stress testing.

[OR], 6.8; 95% CI, 1.3-31.0), left ventricular hypertrophy (OR, 2.1; 95% CI, 1.0-4.5), and any left ventricular systolic dysfunction (OR, 2.0; 95% CI, 1.0-4.1) were independently associated with cardiac events (myocardial infarction, pulmonary edema, ventricular fibrillation or cardiac arrest, and complete heart block in 44 of 570 patients [7.7%]; the absolute event rates corresponding to the ORs were not available).[32] A preoperative echocardiogram with any degree of systolic dysfunction, moderate to severe left ventricular hypertrophy, moderate to severe mitral regurgitation, or an aortic gradient of 20 mm Hg or greater was 80% sensitive for perioperative cardiac events and had a negative predictive value of 97%.[32] A risk model including echocardiographic parameters was more closely associated with perioperative complications than the model including only clinical variables.[32] In other studies, aortic stenosis was associated with increased perioperative death or myocardial infarction (14% vs 2% without aortic stenosis; adjusted OR, 5.2 [95% CI, 1.6-17.0]; $P < .001$).[33] Left ventricular ejection fraction less than 30% was associated with a greater risk of perioperative death, myocardial infarction, and heart failure exacerbation (53.6% vs 26.0% with left ventricular ejection fraction ≥30%; adjusted OR, 4.9 [95% CI, 1.8-14.4]; $P = .008$).[34] However, in a study of 339 men with suspected CAD, echocardiographic measurements did not contribute to the clinical factors associated with perioperative risk.

It is reasonable to consider preoperative echocardiography in patients with moderate or severe valvular disease (stenosis or regurgitation) without echocardiography within the past year, or in those who have new clinical signs or symptoms of severe valvular disease, including dyspnea, angina, edema, or recent syncope.[16]

Treatment of severe valvular disease should be considered before noncardiac surgery. Patients with established or suspected hypertrophic cardiomyopathy, in whom hyperdynamic ventricular function may lead to systolic anterior motion of the mitral valve and left ventricular outflow tract obstruction, and high-risk patients undergoing cardiac solid-organ transplantation may benefit from preoperative echocardiographic evaluation.[35-37] Overall, except in special circumstances, routine preoperative evaluation of ventricular function is not recommended.[16,18,32,38,39]

## Assessment of Functional Capacity and Stress Testing for Myocardial Ischemia

Poor functional capacity is associated with increased risk of perioperative complications.[15] Patients unable to perform workloads of 4 METs or greater, such as walking up a hill or climbing 2 or more flights of stairs, have a 2-fold increased risk of perioperative cardiovascular complications compared with those who are able (9.6% vs 5.2%, respectively; $P = .04$).[15] Among 1396 patients, the quantitative Duke Activity Status Index, derived from a validated questionnaire assessing functional capacity (range, 0-58.2; higher values indicate greater functionality),[40] was independently associated with death or myocardial infarction in 28 patients (2%) within 30 days of surgery (adjusted OR, 0.91 [95% CI, 0.83-0.99]) for every 3.5 points on the index; however, the absolute event rates corresponding to the ORs were not available).[41]

Exercise ECG stress testing assesses functional capacity and identifies stress-induced myocardial ischemia. Pharmacological stress testing is reserved for patients who are unable to exercise. In a study of 530 patients undergoing dobutamine stress echocardi-

ography prior to noncardiac surgery, ischemia at a low workload (<60% of the maximum predicted heart rate) was associated with increased event rates (23% risk of death or myocardial infarction in patients with ischemia vs 5% without ischemia; adjusted, OR, 7.0 [95% CI, 2.8-17.6]).[42] No MACE occurred in patients without preoperative ischemia, whereas 43% of those with ischemia at low workloads had MACE. In a separate study of 429 participants, dobutamine stress echocardiography had an excellent negative predictive value (98%) for perioperative MACE.[43]

Routine stress testing is not indicated for low-risk patients, which includes those with excellent functional capacity (>10 METs, which is equivalent to playing singles tennis or running at a 10 minute/mile pace) and with moderate to good functional capacity (≥4-10 METs, which is equivalent to playing doubles tennis or cross-country hiking). Cardiopulmonary exercise testing may be considered for patients with unknown functional capacity scheduled for higher-risk surgical procedures (Box 3),[16] but it is not recommended by European guidelines.[18] Canadian guidelines recommend against both preoperative exercise stress testing and cardiopulmonary exercise testing due to limited data supporting testing.[39]

Among patients with poor functional capacity (<4 METs) at higher risk for noncardiac surgery, exercise testing with cardiac imaging or noninvasive pharmacological stress testing (either dobutamine stress echocardiography or vasodilator stress myocardial perfusion imaging) to assess for myocardial ischemia is only reasonable if the results from this testing would change perioperative medical management and decisions regarding coronary revascularization.[16,42-44] European guidelines recommend stress testing with imaging before high-risk surgery in patients with more than 2 clinical risk factors (using the Revised Cardiac Risk Index) and poor functional capacity (<4 METs) and treatment according to clinical indications independent of surgery.[18] In contrast, Canadian guidelines recommend against pharmacological stress echocardiography and radionuclide imaging because the predictive discrimination associated with imaging tests has not been adequately compared with those derived from preoperative risk calculators alone.[39]

### Coronary Angiography and Revascularization

Routine preoperative invasive coronary angiography is not recommended before noncardiac surgery.[16] Invasive angiography may be considered in patients with stress tests that indicate myocardial ischemia, but only if the results of angiography would affect perioperative care. The benefit of noninvasive coronary computed tomographic angiography (CCTA) prior to noncardiac surgery is uncertain. In a study of 234 patients undergoing preoperative CCTA, coronary artery diameter stenosis greater than 50% (MACE in 17.2% with obstructive CAD vs 4.3% without obstructive CAD) and multivessel CAD (MACE in 29.7% with multivessel CAD vs 3.7% without multivessel CAD) provided prognostic data in addition to the Revised Cardiac Risk Index.[45] In a meta-analysis of 11 studies evaluating CCTA prior to surgery, severity and extent of CAD were associated with perioperative MACE (specifically, 2.0% in those without CAD; 4.1% in those with nonobstructive CAD; 7.1% in those with 1-vessel obstructive CAD; and 23.1% in those with obstructive multivessel CAD; $P$ < .001).[46] However, CCTA-diagnosed CAD may overestimate risks,[47] and it is not currently

recommended by clinical practice guidelines for risk stratification prior to noncardiac surgery.[39]

Despite the established risks of CAD, routine coronary revascularization prior to surgery does not improve perioperative outcomes. In the Coronary Artery Revascularization Prophylaxis trial,[48] 510 patients with CAD scheduled for vascular surgery were randomly assigned to coronary artery revascularization before surgery or no coronary revascularization. Postoperative myocardial infarction within 30 days (12% in the revascularization group vs 14% in the no revascularization group; $P$ = .37) and long-term mortality at a median follow-up of 2.7 years (22% vs 23%, respectively; $P$ = .92) were not different between the groups; however, patients with left main CAD and reduced left ventricular ejection fraction were excluded from the trial.[48] Based on these data, routine coronary revascularization is not recommended before noncardiac surgery to reduce perioperative MACE.[16]

In contrast, European guidelines suggest that prophylactic coronary revascularization may be considered before high-risk surgery if there is substantial stress-induced ischemia.[18] Although preoperative coronary revascularization may be performed for a compelling indication independent of surgery, such as for those with acute coronary syndrome,[16] performing surgery within 12 months after coronary stent placement is associated with increased perioperative risks.[7,49] Nonetheless, despite current guidelines, invasive coronary angiography before noncardiac surgery is common and preoperative revascularization is performed in 24% of these cases.[50]

### Biomarker Measurement

Preoperative measurement of biomarkers remains an emerging area of investigation for perioperative risk assessment. Serum levels of B-type natriuretic peptide (BNP), a polypeptide released by cardiomyocytes in response to atrial stretch, or the N-terminal pro-BNP (NT-ProBNP) may be associated with perioperative cardiovascular risk. Based on a meta-analysis[51] of individual patient data from 18 prospective observational studies, preoperative BNP levels greater than 92 pg/mL or NT-ProBNP levels greater than 300 pg/mL were associated with increased risk of death or myocardial infarction at 30 days (21.8% in patients with BNP levels >92 pg/mL or NT-ProBNP levels >300 pg/mL vs 4.9% in patients with natriuretic peptides below these levels). Preoperative natriuretic peptide levels were also associated with improved performance of a risk model that included age, Revised Cardiac Risk Index of 3 or greater, vascular surgery, and urgent surgery for the outcome of 30-day perioperative cardiovascular risk, with a net reclassification index of 18%.[51]

In a prospective cohort study of 10 402 patients undergoing noncardiac surgery, preoperative BNP levels between 100 and 200 pg/mL were associated with a 30-day mortality rate of 0.7%; 200 and 1500 pg/mL, 1.4%; and greater than 1500 pg/mL, 4.0% compared with BNP levels less than 100 pg/mL, which were associated with a 30-day mortality rate of 0.3%.[52] Canadian guidelines recommend measurement of NT-proBNP or BNP levels prior to noncardiac surgery in patients with cardiovascular disease, a Revised Cardiac Risk Index of 1 or greater, or for those who are aged 65 years or older.[39] The AHA/ACC guidelines do not formally endorse a BNP measurement as part of preoperative risk assessment because biomarker-based perioperative management strategies have not been tested to reduce cardiovascular risk.[16]

Cardiac troponin level, a sensitive marker of myocardial injury, should be measured perioperatively when signs or symptoms suggest myocardial ischemia or myocardial infarction.[16] Routine cardiac troponin screening should be avoided in unselected patients without symptoms of myocardial ischemia.[16] The value of postoperative cardiac troponin surveillance in asymptomatic patients at risk for ischemic complications is uncertain because no studies have evaluated the benefits of a testing strategy.[16] However, Canadian guidelines recommend postoperative cardiac troponin surveillance in high-risk individuals (eTable 2 in the Supplement).[39] In the authors' opinion, postoperative surveillance of cardiac troponin level during the first 48 hours after higher-risk surgery is reasonable to detect silent myocardial injury in patients at increased risk of cardiovascular events based on preoperative risk calculators (eg, Revised Cardiac Risk Index >1) if the results of testing would modify clinical management (eg, initiation or intensification of antithrombotic or statin therapy for the prevention of cardiovascular events).

## Medical Therapies to Reduce Perioperative Cardiovascular Risk

### β-Blockers

Perioperative use of β-blockers confers a number of potentially advantageous effects on perioperative risk. The use of β-blockers decreases myocardial wall stress, prolongs coronary diastolic filling time, and reduces mismatch in myocardial oxygen supply and demand. Despite observational data suggesting an association of perioperative use of β-blockers with improved outcomes in high-risk patients,[53,54] randomized clinical trial results do not support perioperative prescription of β-blockers.[55] In the Perioperative Ischemic Evaluation trial, 8351 patients were randomly assigned to extended-release metoprolol succinate (100 mg/d) or placebo, beginning within 4 hours prior to noncardiac surgery and continuing for 30 days. The participants randomly assigned to metoprolol had fewer perioperative nonfatal cardiovascular events (myocardial infarction, cardiac arrest, and cardiovascular death; 5.8% vs 6.9% among those assigned to placebo; P = .04), but increased rates of perioperative stroke (1.0% vs 0.5%, respectively; P = .005) and all-cause mortality (3.1% vs 2.3%; P = .03).[56]

It is possible that the longer duration of β-blocker administration prior to surgery and lower doses (or titration to heart rate) may be beneficial. In an observational analysis of 940 patients undergoing vascular surgery, fewer cardiovascular events (myocardial infarction or injury, stroke, or mortality) occurred when β-blockers were initiated more than 1 week prior to surgery compared with shorter preoperative durations (15% vs 27%, respectively; P < .001).[57] Patients already taking β-blockers should continue treatment during the perioperative period in the absence of bradycardia or hypotension.[16] Initiation of β-blockers before surgery may be warranted in select patients with CAD or with multiple risk factors and at high risk for perioperative myocardial infarction.[16] Although high-dose β-blocker therapy should not be initiated on the day of surgery, it may be reasonable to initiate β-blocker therapy more than 1 week prior to surgery to determine tolerability and safety.

### Aspirin

Competing risks of bleeding and thrombosis represent a key challenge during the perioperative period. Aspirin, an irreversible inhibitor of cyclooxygenase-1, reduces platelet aggregation and thrombotic risk by diminishing thromboxane A2 production, with associated increased risks of bleeding. The Perioperative Ischemic Evaluation-2 trial tested the use of routine perioperative aspirin vs placebo in 10 010 patients at risk for cardiovascular complications who were scheduled for noncardiac surgery.[58] Patients assigned to preoperative aspirin did not have significantly lower rates of death or myocardial infarction (7.0% vs 7.1% for those assigned to placebo; P = .92), but aspirin was associated with increased rates of major bleeding (4.6% vs 3.8%, respectively; P = .04). In this trial, only one-third of patients had established vascular disease, and less than 5% had prior coronary stent placement. Thus, routine perioperative aspirin use prior to noncardiac surgery is not recommended,[16] although aspirin therapy may be appropriate for certain patients if ischemic risks outweigh the risks of bleeding.

### Lipid-Lowering Therapy and Statins

Observational data and small randomized trials suggest that lipid-lowering therapy may be associated with lower perioperative cardiovascular risk. In a retrospective, propensity-matched analysis[59] of 204 885 patients undergoing noncardiac surgery, prescribing lipid-lowering drugs during the surgical hospitalization was associated with lower in-hospital mortality compared with patients who did not receive lipid-lowering therapy (2.1% vs 3.1%, respectively; adjusted OR, 0.62 [95% CI, 0.58-0.67]). Similar results were reported from a Veterans Affairs patient cohort (in-hospital mortality of 1.8% with lipid-lowering therapy vs 2.3% in those without lipid-lowering therapy; relative risk, 0.82 [95% CI, 0.75-0.89]) and from the Vascular Events in Noncardiac Surgery Patients Cohort Evaluation study.[60,61]

The Lowering the Risk of Operative Complications Using Atorvastatin Loading Dose trial[62] randomly assigned 648 statin-naive patients with cardiovascular disease (approximately 25%) or multiple risk factors (approximately 75%) to high-dose atorvastatin or placebo within 18 hours before noncardiac surgery, and continued treatment for 7 days postoperatively. Atorvastatin did not reduce major cardiovascular complications (16.6% vs 18.7% for those assigned to placebo; hazard ratio, 0.87 [95% CI, 0.60-1.26]; P = .46).[62] Results from meta-analyses of randomized trials are inconsistent.[63,64] Although randomized trials do not support prescribing statins prior to surgery, the AHA/ACC guidelines suggest that preoperative initiation of statin therapy is reasonable prior to vascular surgery, and the authors' opinion is that statin therapy may be beneficial with few adverse effects in patients with indications for lipid-lowering therapy, such as those with diabetes or atherosclerotic cardiovascular disease who are scheduled for higher-risk surgery.[16]

### Angiotensin-Converting Enzyme Inhibitors and Angiotensin Receptor Blockers

The safety of prescribing angiotensin-converting enzyme inhibitors (ACEIs) or angiotensin receptor blockers (ARBs) on the day of surgery is unclear. In a pooled analysis of 3 small randomized trials including 188 participants, perioperative continuation of ACEIs or ARBs was associated with increased rates of intraoperative hypotension (57.8% vs 23.5% in those who discontinued use of ACEIs or ARBs; pooled relative risk, 2.53 [95% CI, 1.08-5.93]).[39,65-67] In a large observational study of 4802 individuals undergoing noncardiac surgery, perioperative discontinuation of ACEIs or ARBs prior to surgery was associated with a lower risk of perioperative hypotension (23.3% vs 28.6% in those with continued use of ACEIs or ARBs;

**Figure 2. Perioperative Risk Assessment and Management of Patients With a Coronary Stent**

| (1) Assess and integrate all risk considerations. | | | |
|---|---|---|---|
| **Timing and characteristics of coronary stent** | | | |
| Time since percutaneous coronary intervention (PCI) | <3 mo | <6 mo | 6-12 mo | >1 y |
| Coronary stent type | Drug-eluting stent | | Bare metal stent |
| Lesion and stent length | Longer | | Shorter |
| Original indication for PCI | Acute coronary syndrome | | Stable coronary artery disease |
| **Patient age and comorbidities** | | | |
| Patient age | Older (≥60 y) | | Younger (<60 y) |
| Heart failure | Yes | | No |
| Kidney function measured by estimated GFR | <30 mL/min | | ≥30 mL/min |
| Hemoglobin level | <10 g/dL | | ≥10 g/dL |
| **Surgical considerations** | | | |
| Cardiovasular (CV) risks of surgery | High-risk surgery | | Low-risk surgery[a] |
| Bleeding risk during surgery | High | | Low |
| Urgency of surgery | Urgent or emergent | | Elective |

Greater risk · Less risk · Least risk

(2) Determine appropriate delay to surgery.

(3) Discontinue P2Y$_{12}$ inhibitors 5-7 days before major noncardiac surgery.[b]

(4) Continue 81 mg/d of aspirin unless surgical bleeding risks are prohibitive.

(5) Optimize perioperative lipid-lowering therapy.

(6) Avoid perioperative hemodynamic perturbations (hypotension, hypertension, tachycardia).

GFR indicates glomerular filtraton rate; MACE, major adverse cardiovascular events; P2Y$_{12}$, group of drugs that includes: clopidogrel, ticlopidine, ticagrelor, prasugrel, and cangrelor.

[a] Expected risk of MACE less than 1%. See also Box 3.

[b] Discontinue clopidogrel and ticagrelor 5 days prior to surgery and discontinue prasugrel 7 days prior to surgery.

adjusted relative risk, 0.80 [95% CI, 0.73-0.88]) and with a lower risk of the composite end point of myocardial injury after noncardiac surgery, stroke, and mortality at 30 days (12.0% vs 12.9%, respectively; adjusted relative risk, 0.82 [95% CI, 0.70-0.96]).[68]

Canadian guidelines recommend discontinuing ACEIs or ARBs for 24 hours prior to noncardiac surgery and resuming ACEI or ARB

therapy on the second postoperative day when the patient is hemodynamically stable.[39] European guidelines recommend considering temporary discontinuation of ACEIs or ARBs prior to surgery when prescribed for hypertension, but recommend continuing ACEIs or ARBs in stable patients with heart failure and left ventricular systolic dysfunction.[18] In contrast, the AHA/ACC guidelines indicate that it is reasonable to continue ACEI or ARB therapy, and that if these agents are discontinued, they should be restarted as soon as possible after surgery.[16] Additional investigation is warranted to determine the safety of renin-angiotensin system inhibition during the perioperative period.

## Anticoagulation

Oral anticoagulation is frequently indicated for stroke prevention in patients with atrial fibrillation at risk for thromboembolic events and as treatment for patients with venous thromboembolism or valvular heart disease. In patients with atrial fibrillation, anticoagulation is typically interrupted 2 to 5 days prior to noncardiac surgery (based on anticoagulant pharmacokinetics) to reduce the risk of perioperative bleeding. In a trial of 1884 patients with atrial fibrillation randomly assigned to either perioperative bridging therapy with low-molecular-weight heparin (after discontinuing warfarin 5 days before surgery) or placebo, the incidence of arterial thromboembolism was not different between the groups (0.4% vs 0.3% in the placebo group; P = .01 for noninferiority), but bridging anticoagulation was associated with more perioperative bleeding (3.2% vs 1.3%, respectively; P = .005).[69]

In a study of 3640 patients with atrial fibrillation taking a direct oral anticoagulant, stopping use of the oral anticoagulant 1 to 2 days prior to a procedure with a low bleeding risk (eg, eye surgeries or dental procedures) and 2 to 4 days before a procedure with a high bleeding risk (eg, orthopedic surgeries or vascular surgeries) without perioperative bridging therapy was associated with low rates of arterial thromboembolism (0.33%).[70] Based on the available data, perioperative interruption of oral anticoagulation therapy in patients with atrial fibrillation appears safe and perioperative bridging for patients with atrial fibrillation should not be routinely performed. In contrast, patients with mechanical mitral valves and those at risk for thrombotic events with mechanical aortic valves should receive bridging anticoagulation with heparin prior to noncardiac surgery.

## Special Populations

### Older Adults

Adults aged 65 years or older account for 37% of all inpatient surgeries in the US, and older age is associated with increased cardiovascular risk.[16,71,72] In the Perioperative Ischemic Evaluation-2 trial, being aged 75 years or older was independently associated with an increased risk of postoperative myocardial infarction (9.5% for aged ≥75 years vs 4.8% for aged <75 years; adjusted hazard ratio, 1.89 [95% CI, 1.60-2.23]; P < .001).[58] Age-related changes in cardiovascular physiology, including decreased sympathetic responses to stress, reduced vascular compliance, and impaired baroreceptor responses can lead to labile blood pressure and pulse and enhance susceptibility to perioperative hypotension in older adults.[73,74] Cardiac diastolic dysfunction predisposes to heart failure with small increases in intravascular volume. Aortic stenosis affects 4% of individuals aged 70 to 79 years and 10% of those aged 80 to 89 years,[75] and is associated with higher perioperative risks. Noncardiovascular surgical complications such as infection, respiratory failure, and acute kidney injury are more common in older adults compared with

younger adults (any complication in 26.1% [≥80 years] vs 15.1% [<80 years]; P < .001).[74] In a cohort of 30 254 adults aged 65 years or older undergoing noncardiac surgery, 12.1% developed postoperative delirium, 42.9% experienced functional decline (independent vs partially dependent vs totally dependent), and 29.7% required a new postoperative mobility aid.[76]

Compared with younger individuals, less is known about optimal perioperative care of older adults. Older patients are underrepresented in clinical trials and the guidelines provide few cardiovascular care recommendations for this population.[77,78] General principles of perioperative risk stratification should be followed, with an emphasis on assessing baseline functional impairment in older adults.[78,79] Surgical risk prediction models exist for older adults but are not yet widely used. For example, the Geriatric-Sensitive Perioperative Cardiac Risk Index (Table) was recently developed.[24] Additional studies incorporating cognitive function, frailty, and functional status, which are important components of perioperative cardiovascular risk assessment and outcomes in older adults, are needed.[80]

### Patients Requiring Urgent or Emergency Surgery

Urgent (required within 6-24 hours) or emergency (required within 6 hours) noncardiac surgeries are independently associated with increased risk of surgical morbidity (13.8% for emergency, 12.3% for urgent, and 6.7% for elective) and mortality (3.7% for emergency, 2.3% for urgent, and 0.4% for elective).[81] Preoperative cardiovascular evaluation must consider the benefits of surgery and also alternatives to surgery in the context of cardiovascular risks. When emergency surgery is lifesaving, a thorough cardiovascular risk assessment may not be possible, particularly if it would be unlikely to affect management. Guideline-recommended cardiovascular evaluation prior to urgent surgery may be appropriate to exclude acute cardiovascular conditions that are contraindications to noncardiac surgery (Box 2). If warranted, involvement of cardiovascular anesthesia specialists and careful intraoperative and postoperative hemodynamic monitoring should be considered. Efforts to avoid perioperative tachycardia, hypertension, hypotension, and anemia are prudent.

### Patients With Prior Coronary Stents

Despite recommendations to delay noncardiac surgery after percutaneous coronary intervention (PCI),[16] 3.5% of patients or more undergo noncardiac surgery within 6 months of stent placement.[82,83] Individuals requiring surgery within 1 year after PCI are at increased risk of perioperative events compared with those without coronary stents (8.9% vs 1.5%, respectively; adjusted OR, 2.6 [95% CI, 1.4-4.9]; P < .001).[7] Ischemic risks are inversely related to the length of time between stent placement and noncardiac surgery,[7,49,84,85] and are directly related to prothrombotic surgical trauma and early discontinuation of dual antiplatelet therapy (Figure 2).[49,85-89] In some cases, clinically significant perioperative stent thrombosis and myocardial infarction can occur.[90]

Patients undergoing coronary stent placement should have surgery delayed until the risks associated with delaying surgery outweigh the thrombotic risks of stopping dual antiplatelet therapy. A Veterans Affairs study of 28 029 patients undergoing 41 989 surgeries within 24 months of PCI reported MACE in 11.6% of surgeries performed within 6 weeks of PCI; in 6.4% of surgeries performed between 6 weeks and 6 months; in 4.2% of surgeries performed between 6 months and 1 year; and in 3.5% of surgeries performed beyond 1 year after PCI. Elective noncardiac surgery should be delayed for at least 2 weeks after balloon angioplasty, 30 days after bare metal stent implantation, and 12 months after drug-eluting stent placement, although evidence suggests that surgery 3 to 6 months after drug-eluting stent PCI or longer may be safe.[16,49,83,91] Elective noncardiac surgery after drug-eluting stent PCI may be considered after 6 months or longer if the risk of further delay is greater than the expected risks of myocardial infarction and stent thrombosis.[16] Shorter delays to surgery after PCI require further study.[91]

After coronary stent placement, continuation of single antiplatelet therapy with aspirin is recommended in the AHA/ACC guidelines,[16] whereas European guidelines favor individualized decisions based on bleeding and thrombotic risks.[18] A post hoc subgroup analysis from the Perioperative Ischemic Evaluation-2 study among 470 patients undergoing noncardiac surgery with a prior coronary stent suggests perioperative aspirin use is associated with a reduction in 30-day death or nonfatal myocardial infarction (6.0% vs 11.5% without aspirin use; hazard ratio, 0.50 [95% CI, 0.26-0.95]).[92] Other factors associated with perioperative risks after coronary stent placement include longer lengths of the treated coronary lesion and a history of acute coronary syndrome as the initial indication for stent placement (Figure 2).[88,89]

### Limitations

This review has some limitations. First, a separate systematic literature search was not performed for each subcategory discussed. Therefore, some relevant studies may have been missed.

Second, perioperative care guideline recommendations are limited by the quality and availability of evidence and often rely on expert opinion.

## Conclusions

Comprehensive history, physical examination, and assessment of functional capacity during daily life should be performed prior to noncardiac surgery to assess cardiovascular risk. Cardiovascular testing is rarely indicated in patients with a low risk of major adverse cardiovascular events, but may be useful in patients with poor functional capacity (<4 metabolic equivalent tasks) undergoing high-risk surgery if test results would change therapy independent of the planned surgery. Perioperative medical therapy should be prescribed based on patient-specific risk.

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