control heart rate and reduce blood pressure. Current guidelines recommend reducing systolic blood pressure to 120 mm Hg or less in the first hour in patients with aortic dissection. Intravenous  $\beta$ -blockers are first-line treatment; for hypertension that does not completely respond to  $\beta$ -blockers, intravenous vasodilators (nitroprusside, nicardipine) should be administered. Pain control is often necessary and is best accomplished with intravenous opioids.

An algorithm for the management of acute ascending autic dissections is shown in **Figure 43**. Referral for emergency surgery should be considered in all patients with acute autic dissection and cardiogenic shock, patients with type A autic dissection, and patients with type A intramural hematoma, given the very high mortality rate associated with these conditions. Decisions regarding concomitant aortic arch properties and/or coronary artery bypass graft surgery or coronary artery reimplantation depend on the anatomy of the aortic dissection, involvement of the aortic valve or branch vessels, and various other patient characteristics.

Patients who present with uncomplicated type B aortic syndromes may be treated with medical therapy initially. TEVAR, when compared with medical therapy, is associated with similar clinical outcomes and improved survival and disease progression measures at 2 years. Patients with type B stortic dissection and refractory chest/back pain or hypertension, rapid aortic expansion, or organ malperfusion should undergo aortic repair.

In patients with an intramural aortic hematoma or penetrating aortic ulcer, treatment choices depend on the location of the hematoma or ulcer, progression to aortic dissection, and evidence of aortic enlargement. Immediate aortic repair is indicated in patients with an ascending aortic intramural hematoma or penetrating aortic ulcer and in those with enlargement or progression of disease after detection.

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- Acute aortic dissection classically presents with the sudden onset of ripping or tearing pain in the chest, back, or abdomen.
- Clinical examination findings that increase the index of suspicion for an acute aortic syndrome include pulsus paradoxus, asymmetric blood pressures in the upper extremities, and asymmetric pulses.
- CT angiography is the primary diagnostic test in patients in whom acute thoracic aortic disease is suspected.
- Intravenous β-blockers are first-line treatment to reduce heart rate and control blood pressure in patients with acute aortic dissection without cardiogenic shock.
- Emergency surgery should be considered in all patients with acute aortic dissection and cardiogenic shock, type A aortic dissection, and type A intramural hematoma.

# Role of Genetic Testing and Family Screening

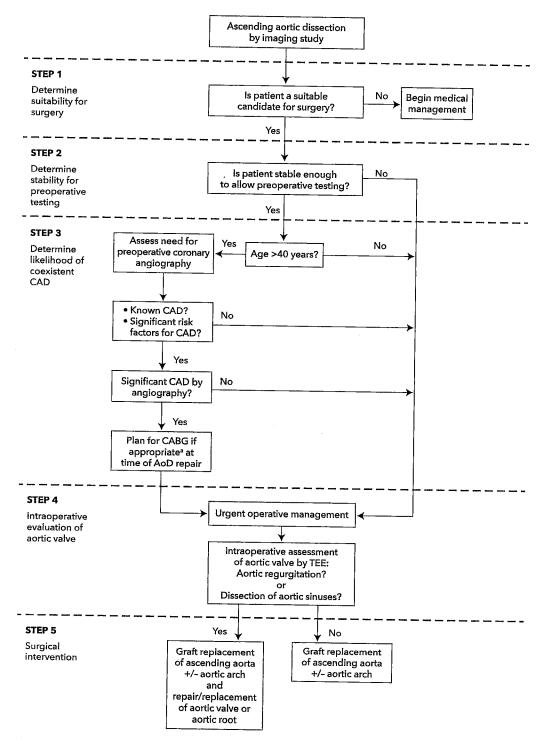
Genetic conditions that predispose patients to thoracic aortic aneurysm syndromes include Marfan, Ehlers-Danlos, and Loeys-Dietz syndromes (see Table 33). Clinical findings and a family history of connective tissue disease often lead to genetic testing, for either diagnostic confirmation or screening of family members. Noninvasive imaging of the aorta should be performed if a pathogenic genetic mutation is found, and routine surveillance (initially at 6 months and then annually, if findings are stable) should follow to ensure that the aorta is not rapidly enlarging.

In first-degree relatives of patients with TAA and/or dissection, noninvasive imaging of the aorta should be performed to identify those with asymptomatic disease. In first-degree relatives of patients with a mutant gene (*FBN1*, *TGFBR1*, *TGFBR2*, *COL3A1*, *MYH11*) and aortic aneurysm and/or dissection, genetic counseling and testing should be performed. Relatives with the genetic mutation should then undergo noninvasive imaging of the aorta.

# **Peripheral Artery Disease Epidemiology and Screening**

Peripheral artery disease (PAD) is most commonly characterized by narrowing of the aortic bifurcation and arteries of the lower extremities, including the iliac, femoral, popliteal, and tibial arteries. Stenosis in the upper extremity arteries, typically at the origin of the subclavian arteries or at branch points of other major vessels, can also occur. Atherosclerosis is the most common cause. Risk factors for PAD include smoking (current or past), diabetes mellitus, and increasing age. PAD occurs at a later age in women than in men, and because women have a longer lifespan, the overall prevalence of PAD is higher in women. The incidence of PAD begins to increase around age 40 years and rises to approximately 10% at age 70 years.

PAD is considered a coronary heart disease risk equivalent, and both asymptomatic and symptomatic patients with PAD are at increased risk for ischemic events, including myocardial infarction, stroke, and cardiovascular death. Patients with atherosclerotic risk factors (smoking, diabetes, hypertension, dyslipidemia, advanced age) who have atypical limb symptoms (leg weakness, paresthesias), exertional leg discomfort, and/or nonhealing ulcers should undergo initial testing with ankle-brachial index (ABI) measurement. According to guidelines from the American Heart Association and American College of Cardiology, a screening ABI is reasonable in asymptomatic persons with one of the following characteristics that signify increased risk: (1) age 65 years and older, (2) age 50 to 64 years with risk factors for atherosclerosis or family history of PAD, (3) age younger than 50 years with diabetes and one additional risk factor for atherosclerosis, or (4) known atherosclerotic disease in another vascular bed (coronary, carotid, subclavian, renal, or mesenteric artery stenosis, or abdominal aortic aneurysm). The U.S. Preventive Services Task Force



**FIGURE 43.** Acute surgical management pathway for AoD. AoD = aortic dissection; CABG = coronary artery bypass graft surgery; CAD = coronary artery disease; TEE = transesophageal echocardiography.

\*Addition of "if appropriate" based on Patel MR, Dehmer GJ, Hirshfeld JW, Smith PK, Spertus JA; American College of Cardiology Foundation Appropriateness Criteria Task Force. ACCF/SCAI/STS/AATS/AHA/ASNC 2009 appropriateness criteria for coronary revascularization: a report by the American College of Cardiology Foundation Appropriateness Criteria Task Force. Force, Society for Cardiovascular Angiography and Interventions, Society of Thoracic Surgeons, American Association for Thoracic Surgery, American Heart Association, and the American Society of Nuclear Cardiology Endorsed by the American Society of Echocardiography, the Heart Failure Society of America, and the Society of Cardiovascular Computed Tomography. J Am Coll Cardiol. 2009;53:530-53. [PMID: 19195618] doi:10.1016/j.jacc.2008.10.005.

Reproduced with permission from Hiratzka LF, Bakris GL, Beckman JA, Bersin RM, Carr VF, Casey DE Jr, et al; American College of Cardiology Foundation/American Heart Association Task Force on Practice Guidelines. 2010 ACCF/AHA/AATS/ACR/ASA/SCA/SCA/SIR/STS/SVM guidelines for the diagnosis and management of patients with thoracic aortic disease A report of the American College of Cardiology Foundation/American Heart Association Task Force on Practice Guidelines, American Association for Thoracic Surgery, American College of Radiology, American Stroke Association, Society of Cardiovascular Anesthesiologists, Society for Cardiovascular Angiography and Interventions, Society of Interventional Radiology, Society of Thoracic Surgeons, and Society for Vascular Medicine. J Am Coll Cardiol. 2010;55:e27-e129. [PMID: 20359588] doi:10.1016/j.jacc.2010.02.015. Copyright 2010, Elsevier.

concluded that there is insufficient evidence to support screening for lower extremity PAD with an ABI in all patients, especially those without risk factors for atherosclerotic vascular disease. Measurement of bilateral arm pressures is indicated in asymptomatic and symptomatic patients with atherosclerotic risk factors to assess for upper extremity PAD.

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- The primary risk factors for peripheral artery disease include cigarette smoking, diabetes mellitus, and advanced age.
- Patients with atypical leg symptoms, exertional leg symptoms, and nonhealing ulcers should undergo ankle-brachial index testing.
- Screening for peripheral artery disease (PAD) with anklebrachial index testing may be considered in patients at increased risk for PAD; evidence does not support screening asymptomatic patients without risk factors.

#### Clinical Presentation

Biscause lower extremity PAD is defined by an abnormal ABI value rather than by symptoms, there is a wide spectrum of clinical manifestations. Patients may present with exertional leg pain relieved by rest (intermittent claudication), atypical exertional leg pain, rest pain, nonhealing wounds, ischemic ulcers, or gangrene. Approximately 25% to 30% of patients with lower extremity PAD present with intermittent claudication, and less than 5% of patients present with critical limb ischemia. Most patients with upper extremity PAD have no symptoms, although patients may present with arm claudication, arm ischemia, or dizziness with arm activity (subclavian steal syndrome).

Patients with intermittent claudication often have reduced exercise capacity and functional status compared with age—and sex-matched controls. Their annual risk for myocardial infarction, stroke, or cardiovascular death is approximately 5% to 7%. Most patients with intermittent claudication have stable symptoms; however, symptoms worsen in approximately 25% of patients, and 10% to 20% of patients will undergo lower extremity revascularization procedures over a period of 5 years.

Critical limb ischemia, the most severe form of PAD, manifests as ischemic rest pain, tissue ulceration, and gangrene. Patients with critical limb ischemia often have reduced exercise capacity and functional status, and these patients have a 30% rate of major amputation and 20% mortality rate within 1 year of diagnosis.

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- Patients with peripheral artery disease may be asymptomatic or present with intermittent claudication, atypical exertional leg pain, or, rarely, critical limb ischemia.
- Patients with peripheral artery disease have reduced
  exercise capacity and an elevated risk for cardiovascular
  morbidity and mortality.

#### **Evaluation**

#### **History and Physical Examination**

A detailed history, review of symptoms, and physical examination are critical in the evaluation of patients suspected of having vascular disease. Patients should be asked about walking impairment, atypical limb symptoms, intermittent claudication, and ischemic rest pain. In patients with exertional leg symptoms, intermittent claudication should be differentiated from pseudoclaudication (symptoms that arise from spinal stenosis) (Table 36). Patients should be questioned about skin breakdown and foot ulcers, and clinicians should educate patients on the importance of foot protection and wearing shoes (specifically, hard-soled shoes) when walking outside the home.

Elements of the physical examination of patients suspected of having PAD are listed in Table 37. Vascular examination of patients suspected of having lower extremity PAD should include comprehensive pulse examination, auscultation for bruits, and inspection of the feet for skin and toenail changes. Patients with PAD may exhibit diminished, absent, or asymmetric pulses, and bruits may be heard at or near sites of arterial stenosis. Patients with critical limb ischemia may have decreased temperature or lack of hair growth in the affected extremity, as well as evidence of poor wound healing or active ulceration (typically involving the digits, plantar aspect of the foot, or heel). Clinicians should distinguish between signs of chronic venous disease (leg edema; pigmented, brawny induration of the gaiter zone; ulceration of the shin or ankle) and critical limb ischemia when evaluating patients with leg ulcers because venous leg ulcers are treated differently than ulcerations in patients with critical limb ischemia (see MKSAP 18 Dermatology).

In patients with upper extremity PAD, a characteristic finding on physical examination is a difference in systolic

TABLE 36. Discriminating Claudication from Pseudoclaudication			
Characteristic	Claudication	<b>Pseudoclaudication</b>	
Nature of discomfort	Cramping, tightness, aching, fatigue	Same as for claudication plus tingling, burning, numbness, weakness	
Location of discomfort	Buttock, hip, thigh, calf, foot	Same as for claudication; most often bilateral	
Exercise-induced	Yes	Variable	
Walking distance at onset of symptoms	Consistent	Variable	
Discomfort occurs with standing still	No	Yes	
Action for relief	Stand or sit	Sit, flexion at the waist	
Time to relief	<5 min	≤30 min	

## TABLE 37. Physical Examination of Patients for Peripheral Artery Disease

Measure blood pressure in both arms (systolic blood pressure difference >15 mm Hg suggests subclavian stenosis)

Auscultate for presence of arterial bruits (e.g., femoral artery)

Palpate for presence of abdominal aortic aneurysm

Palpate and record pulses (radial, brachial, carotid, femoral, popliteal, posterior tibial, dorsalis pedis)

Evaluate for elevation pallor and dependent rubor

Inspect feet for ulcers, fissures, calluses, tinea, and tendinous xanthoma; evaluate overall skin care

blood pressures between the arms, typically more than 15 mm Hg.

#### **Diagnostic Testing**

The most frequently used diagnostic modality to identify lower extremity PAD is measurement of the ABI, which is the ratio of lower extremity to upper extremity systolic blood pressures. ABI measurement is simple, inexpensive, and noninvasive, with a sensitivity and specificity approaching 90%. When undergoing ABI testing, patients should rest for 10 minutes in a supine position before the clinician measures the ankle pressures and brachial pressures with a Doppler machine. Blood pressures should be measured in both arms and in both legs at the dorsalis pedis and posterior tibial ankle locations. To calculate the ABI for each leg, the higher ankle pressure in each leg is divided by the higher brachial artery pressure. In healthy persons, the ankle pressure should be the same as or slightly higher than the brachial pressure; therefore, a normal resting ABI is between 1.00 and 1.40 (Table 38). In the presence of atherosclerotic narrowing of the limb arteries, the downstream blood pressure and concomitant ABI value is lower. A resting ABI of 0.90 or less is diagnostic for PAD and correlates with abnormalities seen on imaging of the arterial tree. A resting ABI greater than 1.40 indicates the presence of noncompressible, calcified arteries in the lower extremities and is considered uninterpretable. In patients with an ABI greater than 1.40, a toe-brachial index is used for diagnosis. A toe-brachial index less than 0.70 is diagnostic for PAD.

TABLE 38. Interpretation of the Ankle-Brachial Index		
Ankle-Brachial Index	Interpretation	
>1.40	Noncompressible (calcified) vessel (uninterpretable result)	
1.00-1.40	Normal	
0.91-0.99	Borderline	
0.41-0.90	Mild to moderate PAD	
0.00-0.40	Severe PAD	
PAD = peripheral artery disease		

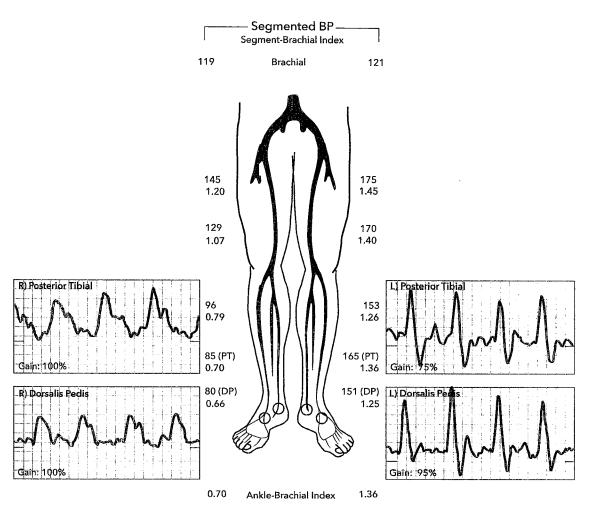
An exercise ABI test is useful when resting ABI values are between 0.91 and 1.40 and the pretest probability of PAD is high. It requires ABI measurements at rest and after treadmill walking or plantar flexion exercises. The American Heart Association has proposed a postexercise ankle pressure decrease of more than 30 mm Hg or a postexercise ABI decrease of more than 20% as a diagnostic criterion for PAD. Other organizations have proposed a postexercise ABI of less than 0.90 and/or a 30-mm Hg drop in ankle pressure after exercise.

Segmental pressure measurements may be performed in a vascular laboratory to localize diseased vessels. This procedure involves pulse volume recordings (measurement of the magnitude and contour of blood pulse volume in the lower extremities) and blood pressure measurements at several locations in the lower extremities (high thigh, low thigh, calf, posterior tibial artery, and dorsalis pedis artery) (Figure 44).

Hemodynamic measurements (ABI, toe-brachial index, exercise ABI, and segmental pressure measurements) remain the most commonly used diagnostic tests for patients suspected of having lower extremity PAD. The ABI does not correlate with the patient's perception of symptom severity or functional limitations; however, lower ABI values are associated with higher rates of myocardial infarction, stroke, and death.

Other imaging tests used to delineate the anatomic location and severity of lower extremity PAD include arterial duplex ultrasonography, CT angiography, and magnetic resonance angiography (Table 39). These imaging modalities are most often used to plan for endovascular or surgical revascularization procedure. Invasive angiography is often preferred because endovascular revascularization procedures can be performed concurrently.

Imaging Modality	Advantages	Limitations
Arterial duplex ultrasonography	Widely available	Limited ability to detect stenoses in the pelvis and in patients with severe calcifications
	Does not require administration of contrast dye	
	Inexpensive	Poor utility for infrapopliteal steno
CT angiography	Widely available	Risk for contrast- induced nephropath
	Useful in defining the severity of PAD	
	Very expensive	
Magnetic resonance angiography	Useful in defining the severity of PAD	Contraindicated in patients with implanted pacemakers and defibrillators
	Very expensive	
		Risk for nephrogen systemic fibrosis in patients with severe kidney disease



**FIGURE 44.** A pulse volume recording demonstrating decreased perfusion in the patient's right side. An ankle-brachial index value of 0.70 on the right side is consistent with the diagnosis of mild to moderate peripheral artery disease (ankle-brachial index <0.90). The normal pulse volume recording on the patient's left side has a sharp systolic upstroke and downstroke (the width of the waveform is much narrower compared with the waveform on the right side). When the amplitude of the waveform is significantly blunted and the width is broader, it suggests moderate to severe disease. BP = blood pressure; DP = dorsalis pedis; PT = posterior tibial.

Duplex ultrasonography, CT angiography, and magnetic resonance angiography are also appropriate in patients with upper extremity PAD to confirm the diagnosis and plan for intervention, such as revascularization.

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- An ankle-brachial index of 0.90 or less indicates peripheral artery disease.
- An ankle-brachial index greater than 1.40 indicates the presence of noncompressible arteries in the lower extremities and is considered uninterpretable; a toe-brachial index is used for diagnosis of peripheral artery disease in these patients.
- Imaging with CT angiography or magnetic resonance angiography is useful to identify the location of stenosis and severity of peripheral artery disease in planning for endovascular or surgical revascularization procedures.

### **Medical Therapy**

Treatment of PAD focuses on reducing cardiovascular risk; improving functional status, quality of life, and claudication symptoms; and preventing tissue loss and amputation.

#### Cardiovascular Risk Reduction

Cigarette smoking is the most important modifiable risk factor for the development of PAD. Smoking cessation is associated with decreased risk for major amputation, improved patency rates following revascularization, and less disease progression. Smoking cessation is imperative to lower the risk for myocardial infarction and stroke and improve overall survival in patients with PAD.

Diabetes is also a strong risk factor for PAD; however, intensive glucose control has not been demonstrated to reduce macrovascular complications, including myocardial infarction, stroke, or amputation. Regardless, patients with PAD and

diabetes should adhere to American Diabetes Association recommendations on diabetes management, with particular attention to foot care.

Dyslipidemia has a mild effect on the development of PAD. Patients with symptomatic PAD should be treated with high-intensity statin therapy, and patients with PAD who are older than 75 years or intolerant of high-intensity statins should be treated with moderate-intensity statin therapy (see MKSAP 18 General Internal Medicine).

Hypertension also has a mild effect on the development of PAD, and control of blood pressure has been associated with reduction of cardiovascular events in patients with PAD. The 2017 high blood pressure guideline from the American College of Cardiology, the American Heart Association, and nine other organizations recommends a blood pressure target of less than 130/80 mm Hg in patients with PAD. There is no consensus on the antihypertensive therapy of choice in patients with PAD; therefore, thiazide diuretics, ACE inhibitors or angiotensin receptor blockers, and calcium channel blockers remain first-line agents in patients with PAD and hypertension. In patients at high risk for cardiovascular events or with comorbid conditions (such as diabetes), ACE inhibitors are effective in reducing cardiovascular morbidity and mortality.

#### Antithrombotic Therapy

Current guidelines recommend antiplatelet monotherapy in patients with PAD to reduce risk for myocardial infarction, stroke, and peripheral arterial events. Despite little supporting evidence, aspirin has been recommended by experts as the primary antiplatelet agent in patients with PAD. In patients who are aspirin intolerant, clopidogrel is recommended as an acceptable alternative. There is no compelling evidence for the use of dual antiplatelet therapy with aspirin plus clopidogrel in patients with PAD alone. In the CHARISMA trial, patients with PAD who were treated with aspirin plus clopidogrel had a reduced rate of hospitalization for myocardial infarction and ischemic events, which was mitigated by a higher rate of bleeding.

Evidence from the TRA 2P-TIMI 50 trial demonstrated that vorapaxar, a thrombin receptor antagonist, was associated with improved limb endpoints (specifically, hospitalization for acute limb ischemia) when compared with placebo in patients with PAD. However, most patients in the TRA 2P-TIMI 50 trial were treated with aspirin, and 30% of patients were treated with aspirin plus clopidogrel. Vorapaxar was also associated with an increased risk for moderate or severe bleeding, including intracranial hemorrhage.

In a large secondary prevention study of patients with prior myocardial infarction receiving aspirin therapy, ticagrelor was superior to placebo in reducing the composite endpoint of cardiovascular death, myocardial infarction, or stroke and major adverse limb events in the subset of patients with PAD. Major bleeding was increased in patients randomly assigned to ticagrelor. However, the absolute risk for major bleeding in patients with PAD was lower than in patients without PAD.

There is no evidence that oral anticoagulation with vitamin K antagonists (such as warfarin) is more effective than antiplately monotherapy in patients with PAD, and anticoagulant therapy's associated with an increased risk for major bleeding. There is also no evidence that non-vitamin K antagonist oral anticoagulants are superior to antiplatelet agents in patients with PAD.

#### **Symptom Relief**

Improving functional status and improving quality of life are critical goals for patients with PAD. In patients who can exercise, supervised exercise training has been associated with improved functional performance and is recommended for patients with intermittent claudication. Systematic reviews comparing supervised exercise with home exercise have reported improvements in maximal walking distance and initial claudication distance that favor supervised exercise; however, no statistically significant differences in quality of life were observed. Despite its effectiveness and safety, supervised exercise training is limited by lack of insurance coverage and unavailability of these programs for patients with claudication.

Medical therapy for patients with intermittent claudication consists of cilostazol, a phosphodiesterase inhibitor with antiplatelet and vasodilator activity. Cilostazol has demonstrated increases in pain-free walking distance and overall walking distance in patients with claudication, and clinical guidelines recommend that patients with claudication be considered for a therapeutic trial of cilostazol. As with other oral phosphodiesterase inhibitors (for example, inotropes such as milrinone), the FDA has placed a black box warning on the use of cilostazol in patients with heart failure. There is no approved pharmacotherapy for patients with critical limb ischemia.

• Smoking cessation is essential to reduce cardiovascular risk in patients with peripheral artery disease.

- Antiplatelet monotherapy with aspirin is recommended for patients with peripheral artery disease to reduce the risk for myocardial infarction, stroke, and peripheral arterial events.
- Supervised exercise training is the most effective treatment for improvement of functional status in patients with peripheral artery disease.
- Cilostazol is recommended for patients with intermittent claudication.

### Interventional Therapy

Endovascular or surgical revascularization procedures are effective in improving symptoms, increasing functional capacity, and improving wound healing in patients with intermittent claudication or critical limb ischemia. Referral for revascularization is indicated in patients with lifestyle-limiting claudication, rest pain, ulceration, or gangrene, especially if there has been an inadequate response to exercise training, cilostazol, and/or wound

treatment. Patients with critical limb ischemia (ABI <0.40, a flat waveform on pulse volume recording, and low or absent pedal flow on duplex ultrasonography) should be considered for urgent revascularization. Endovascular or surgical revascularization should also be considered in patients with a favorable risk-benefit ratio, which is determined by patient factors (age, frailty, comorbid conditions), anatomic factors (severity and burden of atherosclerotic disease, location of disease in lower extremities), operator expertise, and type of procedure. Revascularization is not recommended in asymptomatic patients.

Endovascular revascularization has dramatically increased in recent years because the procedure is minimally invasive and confers a lower risk for perioperative adverse events compared with surgical revascularization. Endovascular revascularization procedures include balloon angioplasty (standard, cutting, drug-coated), stenting (nitinol [nickel-titanium] bare metal and drug eluting), and atherectomy (laser, orbital, rotational, directional). In patients with isolated iliac disease, endovascular revascularization is favored over surgical revascularization owing to lower morbidity and mortality, high procedural success, and high patency rates over time. Most patients undergo balloon angioplasty and stenting of the iliac arteries, given the significantly higher long-term success rate with stenting compared with angioplasty alone.

In patients with femoral, popliteal, or tibial artery (infrainguinal) disease, the patency rates of endovascular evascularization are not as high as in the iliac arteries. Although infrainguinal disease was traditionally treated with augioplasty alone, the advent of atherectomy devices and nitinol stents has changed management. Recently, FDA-approved drug-coated angioplasty balloons and drug-eluting stents have demonstrated superior efficacy compared with standard angioplasty balloons in these patients.

The use of surgical revascularization has declined in the United States; however, patients with complex anatomy that may limit percutaneous procedural success and long-term patency (for example, long chronic total occlusions, multisegment disease) should still be referred for surgical revascularization. The two most common techniques of surgical revascularization are endarterectomy and surgical bypass.

Hybrid revascularization is the concomitant performance of surgical revascularization and endovascular revascularization in a single setting or finite time frame. Hybrid revascularization has increased in conjunction with the rise in endovascular revascularization; however, there are currently no clinical guideline recommendations for hybrid revascularization.

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- In patients with life-limiting claudication who have had an inadequate response to exercise or pharmacologic therapy, endovascular or surgical revascularization is indicated.
- Patients with critical limb ischemia should be considered for urgent revascularization.

#### **Acute Limb Ischemia**



Acute limb ischemia is an infrequent but life-threatening manifestation of PAD. Classically, patients present with at least one of the "6 Ps": paresthesia, pain, pallor, pulselessness, poikilothermia (coolness), and paralysis. Acute limb ischemia is most commonly caused by acute thrombosis of a lower extremity artery, stent, or bypass graft. Other causes include thromboembolism, vessel dissection (usually occurring periprocedurally), or trauma. The presentation of acute limb ischemia represents a true medical emergency; 10% to 15% of patients undergo amputation during initial hospitalization, and 20% of patients die within 1 year.

Anticoagulation, typically with unfractionated heparin, should be initiated as soon as the diagnosis is suspected. Specialists with expertise in revascularization should be consulted, and diagnostic angiography should be performed immediately to define the anatomic level of occlusion. In addition to surgical and endovascular revascularization options, catheter-directed thrombolysis improves outcomes in patients with acute limb ischemia.

Careful monitoring is required after limb reperfusion because of frequent reocclusion, limb edema, and the possibility of compartment syndrome. Signs and symptoms of compartment syndrome include severe pain, hypoesthesia, and leg weakness. If compartment syndrome occurs, surgical fasciotomy is indicated to prevent irreversible neurologic and softtissue damage.

 Acute limb ischemia is characterized by at least one of the "6 Ps": paresthesia, pain, pallor, pulselessness, poikilothermia (coolness), and paralysis.

- Patients with acute limb ischemia should receive immediate anticoagulation therapy and diagnostic angiography in preparation for emergent endovascular or surgical revascularization.
- Careful monitoring after limb reperfusion is required because of frequent reocclusion, limb edema, and the possibility of compartment syndrome.

# Cardiovascular Disease in Cancer Survivors

### Cardiotoxicity of Radiation Therapy to the Thorax

Radiation therapy improves survival in patients with Hodgkin lymphoma, early-stage breast cancer, and other thoracic malignancies. With higher survival rates, cardiovascular disease has emerged as the most common nonmalignant cause of death in patients treated with chest radiation therapy, accounting for 25% of deaths in survivors of Hodgkin lymphoma.