Review article

Surgical management of lower extremity peripheral arterial disease

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Becoming to aging society era, numbers of patients with peripheral arterial disease (PAD) have been continuously increasing. PAD is a presentation of atherosclerotic diseases. Most of the time, the patients are accompanied with underlying diseases that increase risks of atherosclerosis. Treatments of PAD essentially require multidisciplinary team supports. Smoking cessation, controlled exercise program, medical control for underlying diseases must be performed at the beginning of treatments.

Revascularizations are indicated in the patients with unbearable symptoms and limb threatening ischemia. The reason why modern endovascular techniques have expeditiously been grown up is its advantage of less invasive and not to burn the bridge for future surgical bypass. However, its drawbacks are higher reintervention rates, contrast-induced nephropathy, radiation exposure, and lower long-term patency.

Keywords: Peripheral arterial disease, surgical management, lower extremity.

The term peripheral arterial disease (PAD) is classically defined as the various diseases that affect noncardiac, nonintracranial arteries. The most common cause of PAD is atherosclerosis; less common causes include inflammatory disorders of the arterial wall (vasculitis) and noninflammatory arteriopathies, such as fibromuscular dysplasia. In this topic, PAD is confined as a chronic occlusive disease of lower extremities’ arteries due to atherosclerosis. PAD of the lower extremities is the third leading cause of atherosclerotic cardiovascular morbidity, only following coronary artery disease and stroke.

In 2010, estimated 202 million people around the world are living with PAD. PAD usually associated with severe comorbidities especially coronary artery disease (CAD) and cerebrovascular disease (CVD) or both. CAD and CVD are presented in more than half of the patients diagnosed with PAD.

Increasing the prevalence of PAD by age is recognized by a study from the United Nations (UN) in 2010. Of the 40 - 60 year age group, the prevalence of PAD is 3.0 – 10.0% and increases up to 20.0% in the population older than 60 years old. In term of location, half of the PAD lesions occurred in the femoropopliteal artery. Patients with aortoiliac occlusive lesions are usually concomitant with the femoropopliteal disease.

Risk factors

PAD is one of the clinical presentations of systemic atherosclerosis. As such, it shares many risk factors with other clinical atherosclerotic diseases such as CAD and CVD. However, some differences among PAD, CAD, and CVD profiles have been characterized and the strength of associated specific risk factors varies across the clinical syndromes, as well as across anatomic patterns.

Demographic

Age is the strongest risk factor of PAD. The disease is rare among individuals less than 40 years old and rises in prevalence in the sixth to eighth decades. Sex-related differences in PAD appear to be different around the world. Although some studies showed women in some age groups tend to have a lower ankle-brachial index (ABI) than men,
symptomatic PAD still predominates in men. Several studies have demonstrated higher prevalence and poorer outcome of PAD in the black populations compared to the non-Hispanic or Hispanic white populations.

**Smoking**

Cigarette smoking is a strong risk factor of PAD across the population. Several studies revealed an odds ratio in the range of twofold to fourfold. Also, secondhand smoking appears to also carry an increased risk for PAD, although the data in this regard are limited. Smoking cessation is associated with a reduced risk of PAD but it is likely to take more than 20 years of cessation to reduce the level of risk to individuals who have never smoked.\(^{4}\)

**Diabetes**

Diabetes has been strongly related to PAD prevalence and severity. The association is directly related to the severity and duration of diabetes.\(^{5}\) For every 1.0% increase in hemoglobin A1c level, the risk of PAD is increased by 26.0%.\(^{6}\)

**Hypertension**

Hypertension has been associated with most epidemiological studies with an increased risk of PAD, but the relationship has not been as strong as those with smoking or diabetic. Although, systolic blood pressure (SBP) has been associated with PAD, diastolic blood pressure is often not associated with PAD.\(^{7, 8}\)

**Dyslipidemia**

Elevated total cholesterol (TC) and reduced high-density lipoprotein (HDL) levels have been associated with PAD in multiple studies.\(^{7, 9}\) In one study, the TC/HDL ratio has been associated with PAD.\(^{10, 11}\)

**Obesity**

To date, the evidence failed to show a consistent relationship between obesity and PAD.\(^{12}\) The reason for inconsistent is unclear, although it has been postulated that PAD in older people is often associated with other chronic illnesses that might contribute to weight loss.

**Inflammation**

Elevated biomarkers of systemic inflammation have been consistently associated with atherosclerosis in general and PAD specifically. Plasma levels of C-reactive protein and fibrinogen have shown consistent associations with PAD.\(^{11, 13, 14}\) Circulating cytokines such as interleukin-6 and adhesion molecules are also elevated in PAD patients.\(^{15}\)

**Homocysteine**

Some evidence has demonstrated a significant relationship between elevated plasma levels of homocysteine and PAD.\(^{16}\) However, the strength of the association is moderate when controlled for other risk factors.\(^{11}\)

**Socioeconomic status**

PAD has been associated with lower socioeconomic status, including lower levels of income and education.\(^{17, 18}\) These associations span race/ethnicity and are only partly explained by traditional risk factors such as smoking.

**Pathophysiology**

Atherosclerosis is the chronic inflammatory process of the endothelial layer of the blood vessel. After an injury of endothelial cells, the inflammatory process is started with lipid, inflammatory cells such as monocyte, and fibrous material migrates into a wall of blood vessel forming of atherosclerotic plaque. It causes thickening of the blood vessels’ walls and narrowing their lumens. Hence, blood flow to the end organs is reduced. This process is generalized all over the body. In consequence, the clinical syndrome can vary upon which end organs are affected, such as PAD, CAD, or CVD.\(^{19}\) In term of PAD, patients with lower limb ischemia can have the following signs and symptoms, namely:

**Pain**

Pain occurs when blood flow reduction reaches the stage that until oxygen supply does not match oxygen demand. Anaerobic metabolism results in an increase in substance P. Small unmyelinated A-delta and C sensory fibers are stimulated, causing pain. Opposite to oxygen supply reduction, oxygen demand can be increased when patients have activities such as walking or running. Pain on activity is called “intermittent claudication (IC)”, typically should be relieved when activity is stopped. IC can be aggravated at the buttock, thigh, calf or foot regarding the severity and location of atherosclerotic lesions (e.g., infrarenal aorta, iliac arteries, and superficial
femoral artery or lower, respectively). Whenever the disease progresses, the pain could occur even during the resting phase, so-called “ischemic rest pain”. Ischemic rest pain may occur during sleeping while the cardiac output is decreased.

**Tissue loss**

Tissue loss\(^{(20)}\) is a more severe form of PAD, which occurs when ischemia is high severity that makes the necrotic tissue. The two forms are described below, namely:

“Ischemic ulcer” is a result of repeated microtrauma in the ischemic area. A small wound cannot be healed because of inadequate blood flow. Shallow nonhealing pallid erosion is usually found on distal of the foot.

“Ischemic gangrene” mostly occurs on toes when blood flow is insufficient for tissue to survive and results in tissue necrotic. Rutherford category is used to characterize the severity of PAD (Table 1).\(^{(20)}\)

**Clinical presentations**

Asymptomatic patients, the most common form of PAD, have 3 - 4 times more than the symptomatic group.\(^{(21)}\) Asymptomatic patients had no difference in morbidity and mortality from a patient with IC.\(^{(22)}\) Due to underlying diseases, such as congestive heart failure which limits patient activity, diabetic neuropathy which blunts pain sensation, can lead to severe PAD or critical limb ischemia without IC.\(^{(21)}\) Only 7.0% of asymptomatic patients progress to IC. Every 0.1 level dropping of ABI is associated with an increase in the risk of a major cardiovascular event by 10.0 - 13.0%.\(^{(23)}\)

Normally, IC has slow progress. Five years after the onset of symptoms, 70.0 - 80.0% of the patients have stable claudication, only 10.0 - 20.0% will get worsen. Limb amputation will be warranted in 4.0 - 27.0%. Mortality is between 15.0 - 30.0%. Cardiovascular event is a cause of death in half of the patients.\(^{(21, 24)}\)

### Table 1. Rutherford category.\(^{(20)}\)

<table>
<thead>
<tr>
<th>Rutherford category</th>
<th>Clinical description</th>
<th>Objective criteria</th>
</tr>
</thead>
<tbody>
<tr>
<td>0</td>
<td>Asymptomatic—no</td>
<td>Normal treadmill or reactive hyperemia test</td>
</tr>
<tr>
<td></td>
<td>hemodynamically significant occlusive disease</td>
<td></td>
</tr>
<tr>
<td>1</td>
<td>Mild claudication</td>
<td>Completes treadmill exercise(^{†}); AP after exercise &gt; 50 mmHg but at least 20 mmHg lower than resting value</td>
</tr>
<tr>
<td>2</td>
<td>Moderate claudication</td>
<td>Between categories 1 and 3</td>
</tr>
<tr>
<td>3</td>
<td>Severe claudication</td>
<td>Cannot complete standard treadmill exercise(^{†}) and AP after exercise &lt; 50 mmHg</td>
</tr>
<tr>
<td>4</td>
<td>Ischemic rest pain</td>
<td>Resting AP &lt; 40 mmHg, flat or barely pulsatile ankle or metatarsal PVR; TP &lt; 30 mmHg</td>
</tr>
<tr>
<td>5</td>
<td>Minor tissue loss—nonhealing ulcer, focal gangrene with diffuse pedal ischemia</td>
<td>Resting AP &lt; 60 mmHg, ankle or metatarsal PVR flat or barely pulsatile; TP &lt; 40 mmHg</td>
</tr>
<tr>
<td>6</td>
<td>Major tissue loss—extending above TM level, functional foot no longer salvageable</td>
<td>Same as category 5</td>
</tr>
</tbody>
</table>

Categories 4, 5, and 6 are embraced by the term chronic *critical* ischemia.\(^{†}\)

\(5\) minutes at 3 km per hour on a 12.0% incline.

Critical limb ischemia (CLI) is defined when the patients present with ischemic rest pain, ischemic ulcer, or ischemic gangrene. Without intervention, amputation is required in 35.0% of the patients. One-fourth of CLI will die within one year, the most common cause of death is a cardiovascular event.\(^{(25)}\)

**Diagnosis**

The diagnosis of PAD is based on clinical signs and symptoms. History taking and careful physical examination are essential for the diagnosis. Around 20.0 - 50.0% of the patients are asymptomatic.\(^{(3)}\) Ankle-brachial index (ABI) is either a screening or diagnostic tool. The sensitivity and specificity of ABI for diagnosis are 79.0 - 95.0% and more than 95.0%.\(^{(3, 26)}\) ABI less than 0.9 or ankle pressure less than 50 mmHg is a criterion for diagnosis in asymptomatic patients.\(^{(27, 28)}\) The normal range of ABI is between 1.0 - 1.4.

Individuals who have ABI between 0.9 - 1.0 are borderline and require further investigation normally with a treadmill test. Patients would be tested by walking on a treadmill at the speed of 3 km per hour at 12.0% incline for 5 minutes. Normal response is a slight increase or no change in the ABI compared with baseline. If the patient develops no symptom while walking and does not have a corresponding decrease in ankle pressure, PAD is essentially ruling out. Other causes of leg symptoms should be investigated. A falling in ABI more than 20.0% of baseline or dropping ankle pressure below 60 mmHg that requires more than 3 minutes to recover, is considered abnormal.

As for patients who have ABI more than 1.4, the non-compressible vessels are suspected. So, the patients should be organized for further investigation such as toe-brachial pressure index (TBPI) or toe pressure measurement. PAD can be established when TBPI is less than 0.7. Furthermore, Toe pressure less than 30 mmHg indicates advanced ischemia.

**Treatment**

The goal of treatments of lower extremities PAD is quality of life improvement such as slowing disease progression in asymptomatic patients, alleviating claudication, and prevention of limb loss. Lifestyle modification is mandatory in all PAD patients. Atherosclerotic risk modification must be adopted. The target of blood pressure less than 130/80 mmHg, level lipoprotein cholesterol (LDL) level less than 70 mg/dL, and hemoglobin A1C level less than 6.5% are the goals of treatment.\(^{(26, 27)}\)

Time to revascularization depends on the degree of severity of disease. Incapacitating claudication means the claudication causing loss of ability to perform any occupation or basic activity of daily living and critical limb ischemia (CLI) are indications for revascularization.\(^{(21)}\) A recently wound classification is from the Society of Vascular Surgery. Risk stratifications is based on wound, ischemia, and foot infection (WIfI).\(^{(29)}\) Each WIfI category (wound, ischemia, foot infection) is classified into 4 subgroups (grade 0 - 3); then calculates into 4 clinical staging (1 - IV). Each stage carries a different level of risk of amputation and the chances of benefit from revascularization procedure. For instance, patients with a high grade of ischemia but no infection are likely to get benefit from revascularization in contrast to patients with high grade of infection without ischemia have higher chance of amputation and less likely benefit from revascularization.

**Revascularization**

Two different ways of revascularization are open surgery and endovascular therapy. The former is more invasive than the latter but it has fewer reintervention rates. The Bypass versus Angioplasty in Severe Ischemia of the Leg (BASIL) trial\(^{(30)}\), a recent landmark study, comparing open vs. endovascular procedure, showed non-statistically significant difference in early mortality rate (open 5.0% vs. endovascular 3.0%) and more early morbidity in open repair group (57.0% vs. 41.0% in open and endovascular group). These morbidities were mainly infective, wound, and cardiovascular complications. The in-hospital cost of first 12-months follow-up in open-surgery group was estimated as £23,322 (£20,096 of hospital stay and £3,225 of procedure costs), which was about one-third time higher than for those enrolled an endovascular-first group (£17,419 (£15,381, £2,039)).\(^{(30)}\) As an interim analysis, if considering as a whole follow-up period, there was no difference of survival between each group. In addition, for patients who survive longer than two years of randomizations, open surgery group was associated with a significant increase in subsequent overall survival and a trend of improved amputation free survival.\(^{(31)}\)

According to the result of the BASIL trial, open surgical repair is usually recommended in PAD patients, who have estimated life expectancy.
**Surgical risk**

Many pre-operative assessment tools were established such as Revised Cardiac Risk Index (RCRI), American College of Surgeons National Surgical Quality Improvement Program (NSQIP) Myocardial Infarction and Cardiac Arrest (MICA), and Vascular Study Group of New England Cardiac Risk Index (VSG-CRI). As for infrainguinal bypass surgery, VSG-CRI has been proved an accurate tool for prediction, whereas NSQIP-MICA and RCRI provided underpredicted major cardiovascular events.\(^{(32)}\)

The other concern for the endovascular procedure is contrast media exposure. One study showed a significant decline in renal function in patients who were treated with the endovascular intervention compared to those with surgical procedure.\(^{(33)}\)

**Localization of disease**

The localization of disease and extent of occlusion are crucial factors for considering surgical revascularization. TransAtlantic Inter-Society Consensus for management of peripheral arterial disease (TASC-II) divided PAD lesions according to 3 regions; including aorto-iliac (Figure 1), femoral-popliteal (Figure 2), and below knee regions (Figure 3).\(^{(3, 34)}\) Each region is sub-divided into 4 groups, i.e., TASC type A, B, C, and D lesions. Type A is a short or focal stenosis; type D is a chronic total occlusion; type B and C are intermediate lesions.

<table>
<thead>
<tr>
<th>TASC A lesions</th>
</tr>
</thead>
<tbody>
<tr>
<td>• Unilateral or bilateral CIA stenosis</td>
</tr>
<tr>
<td>• Unilateral or bilateral short (≤ 3 cm) EIA stenosis</td>
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</tbody>
</table>

<table>
<thead>
<tr>
<th>TASC B lesions</th>
</tr>
</thead>
<tbody>
<tr>
<td>• Short (≤ 3 cm) stenosis of the infrarenal aorta</td>
</tr>
<tr>
<td>• Unilateral CIA occlusion</td>
</tr>
<tr>
<td>• Single or multiple stenosis totaling 3 to 10 cm involving the EIA not extending into the CFA</td>
</tr>
<tr>
<td>• Unilateral EIA occlusion not involving the origins of internal iliac or CFA</td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>TASC C lesions</th>
</tr>
</thead>
<tbody>
<tr>
<td>• Bilateral CIA occlusions</td>
</tr>
<tr>
<td>• Bilateral EIA stenoses 3 -10 cm long not extending into the CFA</td>
</tr>
<tr>
<td>• Unilateral EIA stenosis extending into the CFA</td>
</tr>
<tr>
<td>• Unilateral EIA occlusion that involves the origins of internal iliac and/or CFA</td>
</tr>
<tr>
<td>• Heavily calcified unilateral EIA occlusion with or without involvement of origins of internal iliac and/or CFA</td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>TASC D lesions</th>
</tr>
</thead>
<tbody>
<tr>
<td>• Infra-renal aortoiliac occlusion</td>
</tr>
<tr>
<td>• Diffuse disease involving the aorta and both iliac arteries requiring treatment</td>
</tr>
<tr>
<td>• Diffuse multiple stenoses involving the unilateral CIA, EIA and CFA</td>
</tr>
<tr>
<td>• Unilateral occlusions of both CIA and EIA</td>
</tr>
<tr>
<td>• Bilateral occlusions of EIA</td>
</tr>
<tr>
<td>• Iliac stenoses in patients with AAA requiring treatment and not amenable to endograft placement or other lesions requiring open aortic or iliac surgery</td>
</tr>
</tbody>
</table>

Figure 1. Inter-Society Consensus for the Management of Peripheral Arterial Disease (TASC) classification of aortoiliac lesions. AAA, abdominal aortic aneurysm; CFA, common femoral artery; CIA, common iliac artery; EIA, external iliac artery.\(^{(35)}\)
TASC A lesions
- Single stenosis ≤ 10 cm in length
- Single occlusion ≤ 5 cm in length

TASC B lesions
- Multiple lesions (stenoses or occlusions), each ≤ 5 cm
- Single stenosis or occlusion ≤ 15 cm not involving the infra geniculate popliteal artery
- Single or multiple lesions in the absence of continuous tibial vessels to improve inflow for a distal bypass
- Heavily calcified occlusion ≤ 5 cm in length
- Single popliteal stenosis

TASC C lesions
- Multiple stenoses or occlusions totaling >15 cm with or without heavy calcification
- Recurrent stenoses or occlusions that need treatment after two endovascular interventions

TASC D lesions
- Chronic total occlusions of CFA or SFA (> 20 cm, involving the popliteal artery)
- Chronic total occlusion of popliteal artery and proximal trifurcation vessels

**Figure 2.** Inter-Society Consensus for the Management of Peripheral Arterial Disease (TASC) classification of femoral-popliteal lesions. CFA, common femoral artery; SFA, superficial femoral artery.\(^3\)

TASC A lesion
- Single focal stenosis, ≤ 5 cm in length, in the target tibial artery with occlusion or stenosis of similar or worst severity in the other tibial arteries.

TASC B lesion
- Multiple stenosis, each ≤ 5 cm in length, or total length ≤ 10 cm or single occlusion ≤3 cm in length, in the target tibial artery with occlusion or stenosis of similar or worst severity in the other tibial arteries.

TASC C lesion
- Multiple stenosis in the target tibial artery and/or single occlusion with total lesion length > 10 cm with occlusion or stenosis of similar or worst severity in the other tibial arteries.

TASC D lesion
- Multiple occlusion involving the target tibial artery with total lesion length > 10 cm or dense lesion calcification or non-visualization of collaterals. The other tibial arteries occluded or dense calcification

**Figure 3.** Inter-Society Consensus for the Management of Peripheral Arterial Disease (TASC) classification of infrapopliteal lesions. The unshaded area represents the target lesion; the area inside the rectangle represents typical background disease.\(^3\)
To date, the patients with TASC type A, B lesions have been recommended for endovascular intervention first while those with TASC type C and D are accompanied with a chance of failure by endovascular revascularization. According to new devices and higher technical (endovascular) skills, angioplasty first strategy or hybrid procedure may be extended to patients with type C and D lesions especially in the patients who are not fit for surgery.\(^{(34, 35)}\)

The American College of Cardiology/American Heart Association (ACC/AHA) guideline\(^{(26)}\) recommends open surgical procedure for patients with acceptable preoperative risk and technical factors advantage over endovascular treatment.

The European Society of Cardiology (ESC) guidelines\(^{(27)}\) recommend as follows. For aortoiliac lesion, open surgery should be considered for a fit patient with an aortic occlusion extending up to the renal artery. Endovascular first is recommended for short occlusive lesion less than 5 cm. As for femoropopliteal occlusive lesions, open surgery is recommended for fit patients with long lesions more than 25 cm of superficial femoral artery lesions when available autologous vein and longer than two year life expectancy.

TASC classification focuses on lesion or involved arterial segment, however, lacking integrating complexity of disease pattern. The global limb anatomic grading system (GLASS) was developed to facilitate decision-making in CLI patient. By incorporating of a baseline assumption, GLASS is less complex and usable in everyday clinical practice and in future research.

GLASS introduced two new concepts, the target arterial path (TAP) and estimated limb-based patency (LBP). The TAP is defined by the treating surgeon or interventionist based on appropriate angiographic image as the optimal arterial pathway to restore in-line (pulsatile) flow to the ankle and foot. LBP is defined as maintenance of in-line flow throughout the TAP, from groin to ankle. The complexity of disease traverse by the TAP is integrated in the GLASS. Femoropopliteal (FP) and infrapopliteal (IP) arterial segments are individually graded on scale of 0 to 4. Using a consensus-based matrix, these grades are combined to three overall GLASS stage (I-III) for the limb.

GLASS includes a simplified approach to the inflow (aortoiliac disease), a dichotomous stratification of severe calcification in the segment, and a simple modifier for pedal (inframalleolar) disease. GLASS stages (I-III) were defined on basis of expected technical success and anatomical durability for infrainguinal endovascular intervention and reflect the overall complexity of disease within the TAP. Inflow disease (aortoiliac and CFA) is considered separately. Furthermore, the inframalleolar disease is not considered within the primary assignment of GLASS staging. In conclusion, GLASS emphasizes on infrainguinal to infrapopliteal disease and is based on some assumptions.\(^{(36)}\)

**Table 2.** Aorto-iliac (inflow) disease staging in GLASS.\(^{(36)}\)

<table>
<thead>
<tr>
<th>Stage</th>
<th>Description</th>
</tr>
</thead>
<tbody>
<tr>
<td>I</td>
<td>Stenosis of the common and/or external iliac artery, chronic total occlusion of either common or external iliac artery (not both), stenosis of the infrarenal aorta; any combination of these</td>
</tr>
<tr>
<td>II</td>
<td>Chronic total occlusion of the aorta; chronic total occlusion of common and external iliac arteries; severe diffuse disease and/or small-caliber (&lt; 6mm) common and external iliac arteries; concomitant aneurysm disease; severe diffuse in-stent restenosis in the AI system</td>
</tr>
</tbody>
</table>

A, no significant CFA disease; B, significant CFA disease (> 50.0% stenosis)
AI, Aortoiliac; CFA, common femoral artery.
A simplified staging system for inflow (AI and CFA) disease is suggested. Hemodynamically significant disease (> 50.0% stenosis) of the CFA is considered a key modifier (A/B)

**Table 3.** Infra-malleolar/Pedal descriptor in GLASS.\(^{(36)}\)

<table>
<thead>
<tr>
<th>Descriptor</th>
<th>Description</th>
</tr>
</thead>
<tbody>
<tr>
<td>P0</td>
<td>Target artery crosses ankle into foot, with intact pedal arch</td>
</tr>
<tr>
<td>P1</td>
<td>Target artery crosses ankle into foot; absent or severely diseased pedal arch</td>
</tr>
<tr>
<td>P2</td>
<td>No target artery crossing ankle into foot</td>
</tr>
</tbody>
</table>
Three GLASS stages were defined based on the likelihood of immediate technical failure (ITF) and 1-year LBP after endovascular intervention of the selected TAP. GLASS stages demonstrate a gradient of infrainguinal disease complexity.\(^{(36)}\)

**Table 4. Femoropopliteal (FP) and infrapopliteal (IP) disease grading in GLASS.**\(^{(36)}\)

<table>
<thead>
<tr>
<th>FP grade</th>
<th>0</th>
<th>1</th>
<th>2</th>
<th>3</th>
<th>4</th>
</tr>
</thead>
<tbody>
<tr>
<td>FP</td>
<td>NA</td>
<td>I</td>
<td>I</td>
<td>II</td>
<td>III</td>
</tr>
<tr>
<td>IP</td>
<td>NA</td>
<td>I</td>
<td>I</td>
<td>II</td>
<td>III</td>
</tr>
</tbody>
</table>

**Table 5. Assignment of GLASS stage.**\(^{(36)}\)

<table>
<thead>
<tr>
<th>Infrainguinal GLASS stage (I-III)</th>
</tr>
</thead>
<tbody>
<tr>
<td>4</td>
</tr>
<tr>
<td>3</td>
</tr>
<tr>
<td>2</td>
</tr>
<tr>
<td>1</td>
</tr>
<tr>
<td>0</td>
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<tr>
<td>0</td>
</tr>
</tbody>
</table>

**Stage I:** low-complexity disease: expected ITF < 10.0% and 1-year LBP > 70.0%

**Stage II:** intermediate-complexity disease: expected ITF < 20.0% and 1-year LBP 50.0 – 70.0%

**Stage III:** high-complexity disease: expected ITF > 20.0% and 1-year LBP < 50.0%

Trifurcation is defined as the termination of popliteal artery at the confluence of the anterior tibial (AT) artery and tibioperoneal (TP) trunk. CFA, common femoral artery; CTO, chronic total occlusion; SFA, superficial femoral artery.

After selection of the target arterial path (TAP), the segmental femoropopliteal (FP) and infrapopliteal (IP) grades are determined from high-quality angiographic images. Using the table, the combination of FP and IP grades is assigned to GLASS stages I to III, which correlate with technical complexity (low, intermediate, and high) of revascularization.
Surgical management of lower extremity peripheral arterial disease

Surgical technique

Endarterectomy

Endarterectomy is a direct removal of obstructive plaque from an arterial segment and it is the best applied for focal lesions in large-caliber vessels, particularly at arterial bifurcations. The advantages of endarterectomy for aorto-iliac lesion are no conduit needed and improving erectile dysfunction.\(^{(37)}\)

After longitudinal arteriotomy was performed, the atherosclerotic plague was bluntly removed from the deep medial layer of the arterial wall. There is a study that showing a 10-years patency rate of endarterectomy of aortoiliac lesions was at 85.7\%.\(^{(38)}\)

With the advent of modern endovascular technology and technique, technical success rates and 5 to 10-year patency rates of endovascular therapy are comparable but less invasive to endarterectomy. These have largely led to an abandonment of endarterectomy in aortoiliac location.\(^{(39)}\)

Arterial bypass

Three factors must have been considered to achieve successful arterial bypass procedures.

- The inflow artery is an artery proximal to the occlusive lesions.
- The outflow artery is a target artery distal to the obstructive lesions.
- Conduit, a bridge or duct, is categorized into an autogenous graft and a synthetic graft. Although an autogenous graft is superior to synthetic in terms of lower patency and infection rates, for the aortoiliac lesion, the vein graft size is too small conduit supplying bilateral outflow arteries of both legs. Polytetrafluoroethelene (PTFE) or polyester woven graft are used for surgical bypass of aortoiliac regions without a difference in patency rates.\(^{(40, 41)}\) One study showed more distal anastomosis stenose in patients with PTFE graft for aortobifemoral bypass.\(^{(42)}\)

Aortobifemoral bypass

Aortobifemoral bypass is indicated in diffuse aortoiliac disease. Transperitoneal, retroperitoneal, and laparoscopic approaches can be considered. Transperitoneal approach advantages in anatomic exposure. Although retroperitoneal approach may cause more difficult to expose of right renal and right iliac arteries, it can keep the patients away from morbidity of intraperitoneal operations such as pain, pulmonary complication, wound complication, ileus, etc. and allows shorter length of hospital stay.\(^{(43, 44)}\)

Laparoscopic aortobifemoral bypass reduces postoperative complications, provides good cost-effectiveness, and comparable patency rate to open surgery but spends longer operative time.\(^{(45)}\) Twenty to thirty cases had usually required for learning curve\(^{(46)}\) and the conversion rate was about 13.0\%.\(^{(43)}\)

The transperitoneal approach starts with long midline incision from xiphoid to the pubis symphysis followed by two longitudinal or oblique incisions at both groins. After inflow and outflow arteries are exposed and controlled, a proximal anastomosis is created. End-to-end anastomosis provides good in-line flow, less turbulence, fewer pseudoaneurysm rates, and better patency rate.\(^{(42)}\) An end-to-side anastomosis is compulsory for those with patent bilateral common iliac and internal iliac arteries but occlusion of bilateral external iliac arteries. Since the end-to-side configuration provides native flow to bilateral internal iliac arteries, contrary to the end-to-end fashion that requires retrograde blood flow through the patent of external iliac arteries for prevention of pelvic organ and buttock ischemia (Figure 4). The next step is tunneling the graft underneath both ureters and performing end-to-side anastomosis from graft to both common femoral arteries or profunda femoral arteries in case of diffuse occlusion of the superficial femoral arteries.\(^{(47)}\) The recommended size of the bifurcated graft is 18 by 9 mm for men and 16 by 8 mm for women.

Five-year and ten-year primary patency rates of aortobifemoral bypass are 85.0 - 90.0\% and 75.0 - 85.0\% respectively. This configuration has the longest patency compared to other extra-anatomical bypass configurations. Early mortality rates are between 4.0 - 4.4\%. Half of the mortalities are caused by cardiovascular events.\(^{(3, 48)}\)

Axillofemoral bypass

An axillofemoral bypass is an alternative option, in case that aortobifemoral bypass is contraindicated such as active intraabdominal infection or high expected perioperative risk for transabdominal operation.\(^{(49, 50)}\) Since 5-year patency rates of axillofemoral and axillounifemoral bypass are lower than that of aortobifemoral bypass (71.0\% and 51.0\% vs. 85.0 - 90.0\%, respectively).\(^{(3)}\) Axillary inflow selection is considered from a SBP of both arms if the difference between each arm is more than 10 mmHg, the higher one is selected for an inflow artery. If there is minimal difference in SBP, the
selected inflow artery is ipsilateral to the target lower extremity. A proximal anastomosis is usually medial to pectoralis minor muscle at first part of the axillary artery to avoid overstretching of anastomosis when patients abduct the arm. The graft is placed in the subcutaneous layer along with the midaxillary line. The most common selected graft size is 8 mm in diameter or 6-mm diameter in patients with small vessel size. No difference in primary patency between ring-supported and unsupported grafts (51) (Figure 5).

**Femoro-femoral bypass**

Femoro-femoral bypass is indicated in unilateral iliac occlusive disease or performs with an axillofemoral bypass. Regional or local anesthesia can be considered for the procedure. Five-year patency rate is 70.0 - 75.0% (3).

Regarding operative techniques, the graft is placed in a prefascial subcutaneous suprapubic plane. For patient with extremely thick or thin abdominal wall or previous surgery or radiotherapy of suprapubic (hypogastric) region, the graft is placed in a preperitoneal plane. Downside of preperitoneal graft placement is higher risk of bowel or bladder injury. End-to-side anastomosis fashion is applied on both femoral arteries. Inverted C configuration is common. Recommended anastomosis length is about 3 times of the graft diameter. The more distal anastomosis may be warranted in obese patients to reduce the chance of graft kinking.

![Figure 4](image-url)

**Figure 4.** Aortobifemoral bypass; A: abdominal aorta, B: end-to-end proximal anastomosis, C: end-to-side proximal anastomosis.

![Figure 5](image-url)

**Figure 5.** (A) axillofemoral bypass, (B) Infrainguinal bypass; Femoral to PTA bypass CFA, common femoral artery; SFA, superficial femoral artery; TPT, tibioperoneal trunk; ATA anterior tibial artery; PTA posterior tibial artery; peroneal a., peroneal artery. (3)
**Infrainguinal (outflow) disease**

Infrainguinal disease is generally defined as atherosclerosis extends down from the superficial femoral artery. Occlusion of outflow vessels is an indication for open surgical repair to fit patients with available autogenous vein graft.\(^{(3,26,27)}\) Since endovascular treatment in long stenosis or occlusion lesions has a lower patency rate compared with surgical bypass, endovascular first concept can be considered for unfit patients. In the case of diffuse occlusive disease, inflow lesion is the priority for treatment.

Planning of surgical bypass requires consideration of 3 issues: inflow; outflow; and conduit. As previous mentioned, the selected inflow artery, proximal to the stenotic lesions, is either femoral artery or popliteal artery. Outflow artery should supply the arch of foot and length of bypass graft should be as short as possible.

Conduits are detours of blood flow from inflow to outflow artery. Many types of conduits have been used for bypass surgery.

**Autogenous vein**

Evidence showed longer primary patency and fewer complication rates in above-and below-knee femoropopliteal bypass by autogenous vein compared to prosthetic graft.\(^{(52)}\) If available, the autogenous vein is the conduit of choice. The autogenous veins of choices are great saphenous vein, small saphenous vein, and arm vein\(^{(53)}\), respectively.

Selection of a good autogenous vein graft based on preoperative evaluation; duplex venous ultrasonography can offer the vein sizing. Larger than 3mm in diameter is required and venosclerosis should be avoided.\(^{(54)}\)

**Prosthetic conduit**

The advantage of prosthetic conduits is availability. The drawback is its shorter patency and a higher infection rate than autogenous vein graft. Prosthetic conduit is a second choice when autogenous vein graft is not available. Polyester graft has a higher 10-year patency rate than PTFE in the above-knee femoropopliteal bypass but not different in below the knee bypass. PTFE with vein cuff had risen the patency.\(^{(52,55)}\) Composite (spliced vein) graft, prosthetic with a vein graft, has similar patency, better secondary patency, more wound complication, and more reintervention rates compared to PTFE graft with vein cuff.\(^{(56)}\)

**Cryopreserved graft**

Cryopreserved graft is a harvested cadaveric vessel preserved by cryopreservation. Some parts of the epithelial lining of blood vessels might be damaged resulting in delayed thrombosis and false aneurysm. Furthermore, the cost is expensive compared to the prosthetic graft and the patency rate is poor. The cryopreserved graft may be useful in infected prosthetic graft and an autogenous vein is unavailable.\(^{(57)}\)

**Human umbilical vein (HUV)**

Human umbilical vein showed a higher patency rate than PTFE but late aneurysmal change can occur. So, HUV is rare in general practice.

**Graft configuration**

The two technique for autologous vein graft arrangement are reversed vein graft and in situ vein graft. The reversed vein bypass graft is performed by flipping vein graft, anastomosing distal site of the graft to inflow artery, then creating the distal anastomosis. The advantage is no requirement of valvulotomy, but the downside is a size discrepancy between vein graft and arterial inflow.

In situ vein bypass graft requires no fully dissection of vein graft from its bed but connection a proximal great saphenous vein to the femoral artery, then making distal anastomosis between the distal vein to outflow artery. In addition, this configuration requires valvulotomy. Evidence observed shows no difference in patency rates between the two configurations.\(^{(58,59)}\)

**Complication**

**Systemic complications**

Systemic complications in the surgical procedure of patients with PAD can be serious due to the patient’s comorbidities. Major complications such as cardiovascular complications and stroke can be found at 4.7% and 1.7%, respectively.\(^{(60)}\)

**Local complications**

**Wound infection**

Wound infection is an early complication and the most common cause of readmission. As for the lower extremities bypass, the wound infection rate is 4.8 – 17.0%. The risk factors includes hypertension, obesity, chronic kidney disease with long term dialysis, ABI less than 0.35, blood glucose higher than 180 mg/dl, operative time longer than 240 minutes,
blood transfusion more than 3 units, iodinated skin preparation, history of smoking, chronic obstructive pulmonary disease, congestive heart failure, and sepsis within 48 hours postoperatively.\(^{(61,62)}\)

**Graft infection**
Graft infection is a dreadful complication that can lead to devastating results included death or limb loss. In general, the treatment of infected graft should be graft removal with appropriate systemic antibiotic treatment. In the case of early infected autogenous vein graft that is not caused by *Pseudomonas aeruginosa* and no clinical of sepsis may be treated by local wound care and systemic antibiotics.\(^{(63,64)}\)

**Graft thrombosis**
Graft thrombosis can be classified into early (<30 days) and late (>30 days). Retained venous valves, anastomotic problems, inadequate local endarterectomy, clamp defects, or inadequate outflow can cause early graft thrombosis.\(^{(65)}\) Thirty percent of late graft thrombosis that occurred in aortoiliac disease results from atherosclerosis progression at the anastomotic site. Other factors related to graft patency are the length of the bypass graft, the quality of the recipient artery, the extent of runoff to the foot, and the quality of conduit.\(^{(35)}\)

**False aneurysm**
False aneurysm rates are around 1.0 - 5.0% in aortoiliac bypass. There are several causes such as infection, fabric degeneration, and following endarterectomy technique. Most of the time, false aneurysm at aortic anastomosis are discovered by surveillance imaging but pulsatile mass can be palpable at the common femoral anastomosis. The treatment of choice is debridement and open repair with interposition graft.\(^{(66)}\)

**Aortoenteric fistula**
Aortoenteric fistula is a rare but lethal complication. This condition results from end-to-side anastomosis between aorta and graft eroding into the adjacent bowel (most common; third-part of duodenum), causing gastrointestinal bleeding that may be severe and lead to death. Operative correction is warranted. \(^{(67)}\)

**Postoperative follow-up**
Routine surveillance should be performed at 4 to 6 weeks post-operative then every 6 months for the first year, then annually.\(^{(26)}\) History taking, physical examinations, pulse examination at proximal artery, graft, outflow vessel, and ABI measurement are mandatory for surveillance. Symptoms and signs of limb ischemia or decrease ABI more than 0.15 indicates stenosis or occlusion. Surveillance imaging such as duplex ultrasonography (DUS) is more sensitive to detect graft thrombosis than ABI, especially in autogenous vein graft. DUS can estimate the degree of stenosis by calculating peak velocity ratio. Normal peak velocity ratio is less than 2.0. If the peak velocity ratio is more than 3.0, reintervention may be indicated.\(^{(68)}\)

**Conclusion**
PAD results from atherosclerosis and is usually concomitant with several diseases or conditions such as hypertension, diabetes, hyperlipidemia, and smoking, etc. The physician should be suspicious and look for signs and symptoms of PAD in patients with risk factors. ABI is an easy screening and diagnostic tool. The goal of treatment is to increase the quality of life and prevent limb loss. Treatment consists of lifestyle modification, medical control of risk factors, and revascularization if indicated.

The keys to success in revascularization are patient selection, preoperative evaluation, planning, and meticulous surgical bypass. Post-operative long-term surveillance is mandated.

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