# Peripheral Arterial Disease – The New Cardiovascular Endemic

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# **Executive Summary**

- Peripheral artery disease or PAD commonly refers to the presence of a stenosis or occlusion in the aorta or arteries of the limbs
- Individuals with PAD have an exceptionally elevated risk for cardiovascular events, and the majority will eventually die of cardiac or cerebrovascular etiology
- Prognosis is correlated with the severity of PAD as measured by the ankle brachial index (ABI)
- General practitioners (e.g., PCP, podiatrists, etc) must be engaged in the diagnosis and management of PAD—it can be life saving
- Early referral to a (cardio)vascular specialist can facilitate optimal risk factor modification and management—this saves lives
  - WHEN revascularization is necessary, endovascular therapy for PAD should be considered FIRST-LINE therapy in most cases

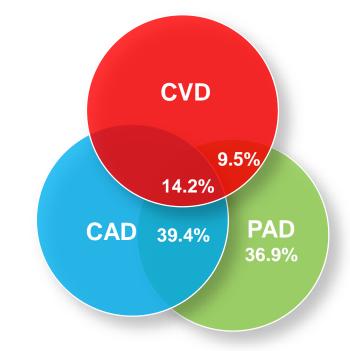


# PAD Epidemiology



# Definition of PAD<sup>1,2</sup>

- The presence of a stenosis or occlusion in the aorta or arteries of the limbs
- One of the three cardinal manifestations of atherosclerosis in addition to CAD and CVD
- Associated with an increased risk of cardiovascular and cerebrovascular events, including death, MI and stroke



Patients with one manifestation often have coexistent disease in other vascular beds<sup>1</sup>

CAD=coronary artery disease; CVD=cardiovascular disease; MI=myocardial infarction.

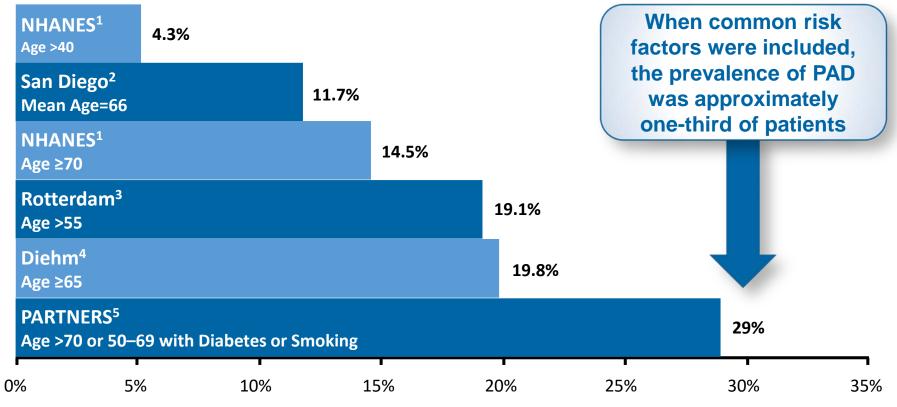
1. Bhatt DL et al, on behalf of the REACH Registry Investigators. JAMA 2006; 295(2): 180-189

2. Rooke T et al. 2011 ACCF/AHA focused update of the guideline for the management of patients with peripheral arterial disease (updating the 2005 guideline): a report of the American College of Cardiology Foundation/American Heart Association Task Force on Practice Guidelines.

Circulation. 2011;124:2020–2045.



### Prevalence of PAD



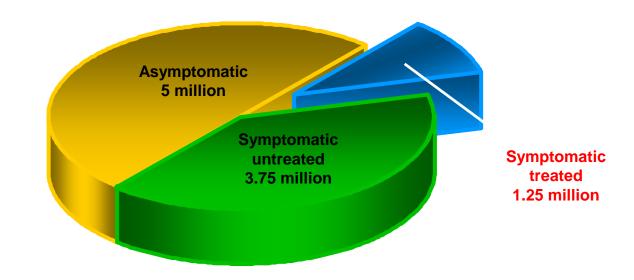
#### NHANES=National Health and Nutrition Examination Survey.

- 1. Selvin E, Erlinger T. Prevalence of and risk factors for peripheral arterial disease in the United States: results from the National Health and Nutrition Examination Survey, 1999–2000. *Circulation*. 2004;110:738–743.
  - 2. Criqui M et al. The prevalence of peripheral arterial disease in a defined population. *Circulation*. 1985;71:510–515.
- 3. Meijer W et al. Peripheral arterial disease in the elderly: the Rotterdam Study. *Arterioscler Thromb Vasc Biol.* 1998;18:185–192.
- 4. Diehm C et al. High prevalence of peripheral arterial disease and co-morbidity in 6880 primary care patients: cross-sectional study. Atherosclerosis. 2004;172:95–105.
  - 5. Hirsch A et al. Peripheral arterial disease detection, awareness, and treatment in primary care. *JAMA*. 2001;286:1317–1324.



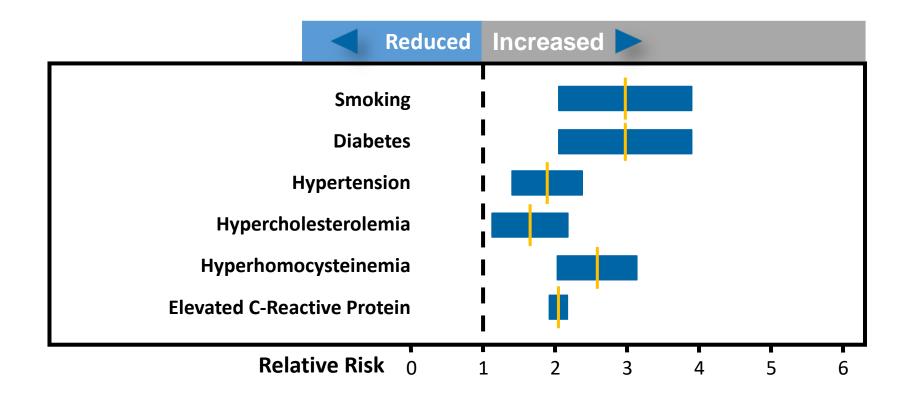
# Prevalence

#### Total ~ 10 million U.S. patients



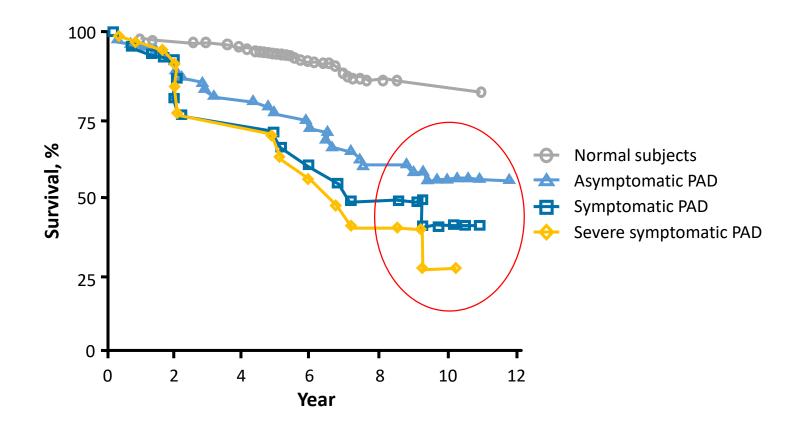


# Risk Factors for PAD



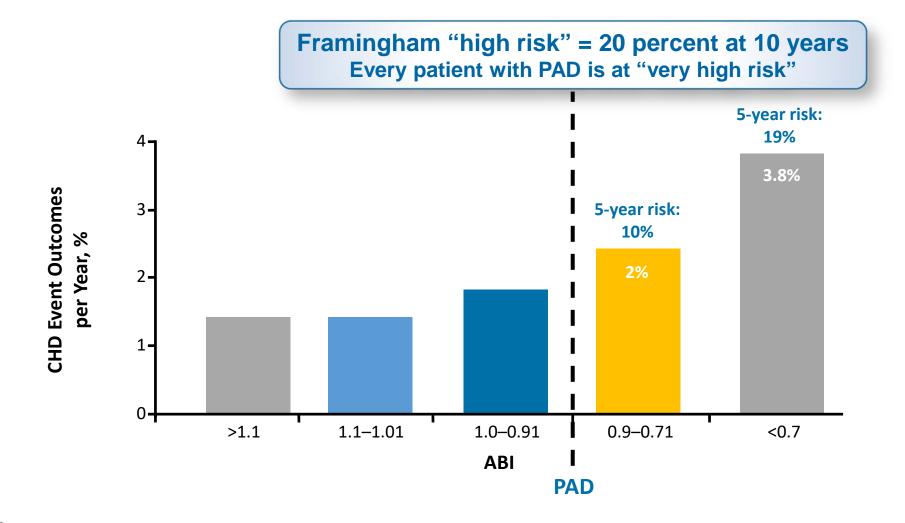


### 10-Year Survival Rates for Patients with PAD





### Cardiovascular Risk Increases with Decreases in ABI





Fatal or nonfatal MI.

CHD=coronary heart disease (chronic heart failure).

## Clinical Presentation

•	Asymtomatic	20-50 %
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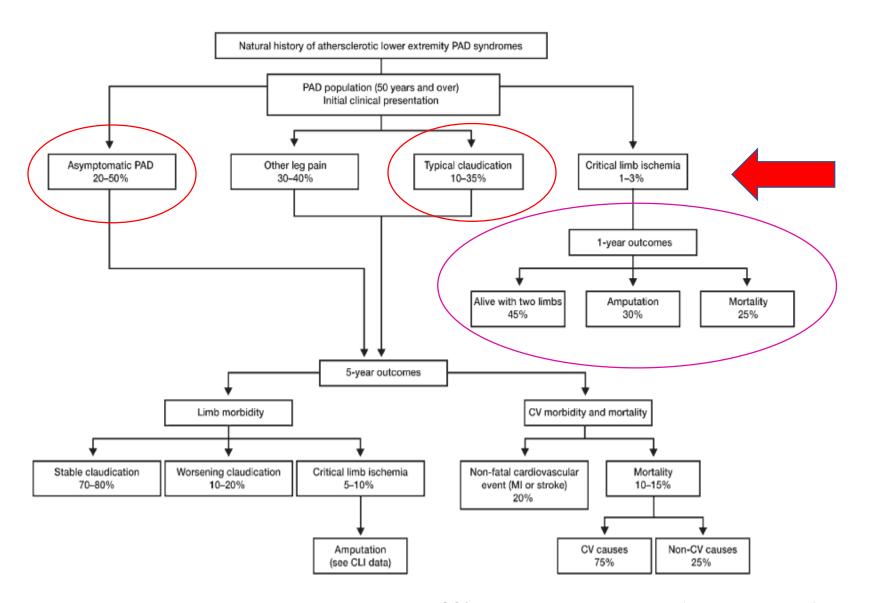
- Atypical leg pain40-50 %
- Classic claudication 10-35 %
- Critical limb ischemia 1-2 %



# Clinical Manifestations of PAD

- Patients can be asymptomatic for PAD (up to 50-60% of the time)
- Or patients can experience
  - Symptoms of intermittent claudication
    - Discomfort
    - Aching
    - Leg cramps with exercise that resolve with rest
  - Functional impairment
    - Slow walking speed, gait disorder
  - Rest pain
    - Pain or paresthesia in foot or toes, worsened by leg elevation and improved by dependency
  - Ischemic ulceration and gangrene (Critical Limb Ischemia)







Hirsch, AT et al. ACC/AHA 2005 Practice guidelines for management of patients with PAD, Circ 2006

# Classification of PAD

	Fontaine Stages	Rutherford Categories		
Stage	Clinical	Grade	Category	Clinical
I	Asymptomatic	0	0	Asymptomatic
IIA	Mild claudication	I	1	Mild claudication
IIB	Moderate-severe claudication	I	2	Moderate claudication
IID		I	3	Severe claudication
Ш	Ischemic rest pain	П	4	Ischemic rest pain
IV	Ulceration or gangrene	III	5	Minor tissue loss
IV		IV	6	Ulceration or gangrene



# Diagnosis of PAD





# Comprehensive Vascular Examination

#### **Pulse\* Examination**

- Carotid
- Radial/ulnar
- Femoral
- Popliteal
- Dorsalis pedis
- Posterior tibial

#### **Physical Examination**

- Bilateral arm blood pressure
- Cardiac exam
- Palpation of abdomen for potential aneurysmal disease
- Auscultation for bruits
- Examination of legs and feet



# Physical Examination Findings Suggestive of PAD

- Performed with pants and shoes off
- Limb examination (and comparison with the opposite limb) includes
  - Absent or diminished femoral or pedal pulses (especially after exercising the limb)
  - Arterial bruits
  - Hair loss
  - Poor nail growth (brittle nails)
  - Dry, scaly, atrophic skin
  - Dependent rubor
- Pallor with leg elevation after 1 minute at 60° (normal color should return in 10–15 seconds; >40 seconds indicates severe ischemia)
- Ischemic tissue ulceration (punched-out, painful, with little bleeding), gangrene





# Diagnosis

- ABI with exercise
- Dupplex ultrasound
- CTA/MRA
- Conventional angiography



#### Noninvasive Assessments in CAD and PAD Are Similar

#### CAD

- Electrocardiogram
- Echocardiogram
- Stress test
- Angiography

#### **PAD**

- ABI
- PVR, segmental pressures
- Treadmill test
- DUS, CTA, MRA



Hirsch A et al. ACC/AHA 2005 practice guidelines for the management of patients with peripheral arterial disease (lower extremity, renal, mesenteric, and abdominal aortic): a collaborative report from the American Association for Vascular Surgery/Society for Vascular Surgery, Society for Cardiovascular Angiography and Interventions, Society for Vascular Medicine and Biology, Society of Interventional Radiology, and the ACC/AHA Task Force on Practice Guidelines (writing committee to develop guidelines for the management of patients with peripheral arterial disease): endorsed by the American Association of Cardiovascular and Pulmonary Rehabilitation; National Heart, Lung, and Blood Institute; Society for Vascular Nursing; TransAtlantic Inter-Society Consensus; and Vascular Disease Foundation. Circulation. 2006;113:e463–e654.

### ACCF/AHA 2011 PAD Guidelines Diagnostic Methods: ABI

- The resting ABI should be used to establish the lower extremity PAD diagnosis in patients with suspected lower extremity PAD, defined as individuals
  - With exertional leg symptoms
  - With nonhealing wounds
  - Who are age ≥65
  - Or who are age ≥50 with a history of smoking or diabetes





### Ankle Brachial Index

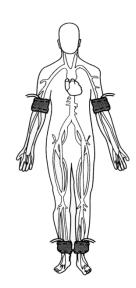
Ankle systolic pressure

ABI =

#### **Brachial systolic pressure**

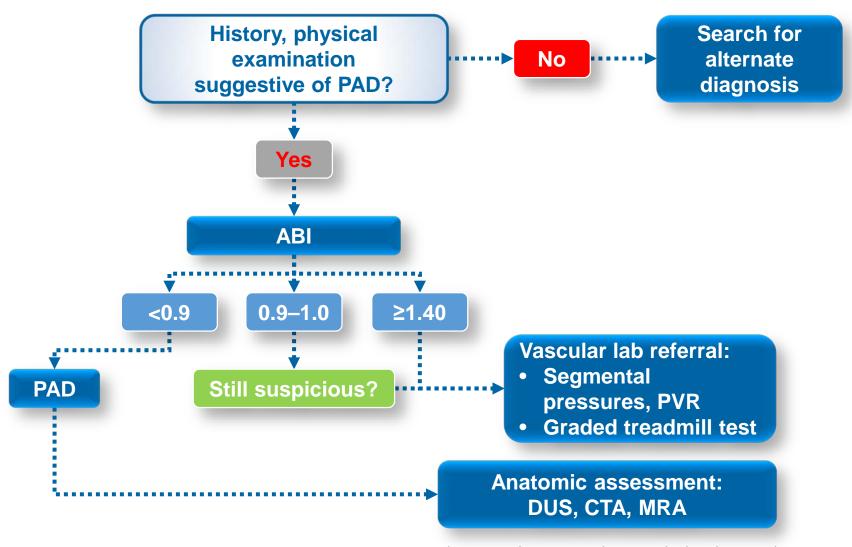
- Ankle and brachial systolic pressures taken using a hand-held Doppler instrument
- The ABI is 95-percent sensitive, 99-percent specific for PAD

Normal	1.00–1.40		
Borderline	0.91–0.99		
PAD	≤0.90		
Pain/Ulceration	≤0.40		
Noncompressible	≥1.40		



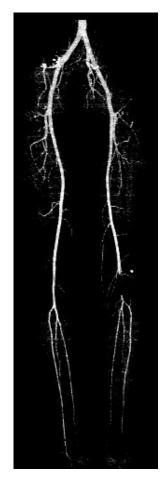


## Suggested Diagnostic Algorithm for PAD





# Advanced Vascular Imaging



# **CT Angiography**

- Maximum-intensity projection (MIPs)
  - Angiographic like representation
- Volume rendering
  - Preserves depth information
- Multi-planar reformat
- Curved planar reformat (CPR)
  - Perpendicular to median arterial centerline



#### **MR** Angiography

- Traditional: Time of flights
- Contrast-enhanced MRA
  - Improves speed of exam, anatomic coverage, and small- vessel resolution
- Time-resolved gadolinium enhanced sequences
  - Time-resolved imaging of contrast kinetics (TRICKS)
  - Provides angiographic like dynamic contrast passage
- Moving-table technique or multi-array, parallelimaging
  - Optimize large field-ofview imaging



# Management

# Medical Therapy and Risk Factor Modification



# Therapies for PAD



#### Preventing death, MI, stroke

- Antiplatelets (Pletal/ASA/Plavix/Zontivty)
- Cholesterol lowering statins
- Angiotensin-converting enzyme (ACE) inhibitors

#### **Reducing symptoms**

- Exercise
- Pletal (Cilostazol).
- Endovascular interventions
- Surgery

#### **Saving limbs**

- Endovascular interventions
- Surgery



# Pharmacotherapy

- Drugs with evidence of clinical utility in claudication
  - Cilostazol [Phosphodiesterase Inhibitor]
- Drugs with supporting evidence of clinical utility in claudication
  - Carnitine and Propionyl L-Carnitine [skeletal muscle oxidative metabolism]
  - Lipid lowering agents
- Drugs with insufficient evidence of clinical utility in claudication
  - Pentoxifylline
  - Antithrombotic agents [ASA/plavix]
  - Vasodilators [CCB/ $\alpha$ -adrenergic antagonists/β2-adrenergic agonists/Papaverine]
  - 5-hydroxytryptamine antagonist [Ketanserin/Sarpogrelate]
  - Prostaglandins [PGE1]



#### Recommendation 14

Exercise therapy in intermittent claudication

- Supervised exercise should be made available as part of the initial treatment for all patients with peripheral arterial disease [A].
- The most effective programs employ treadmill or track walking that is of sufficient intensity to bring on claudication, followed by rest, over the course of a 30-60 minute session. Exercise sessions are typically conducted three times a week for 3 months [A].

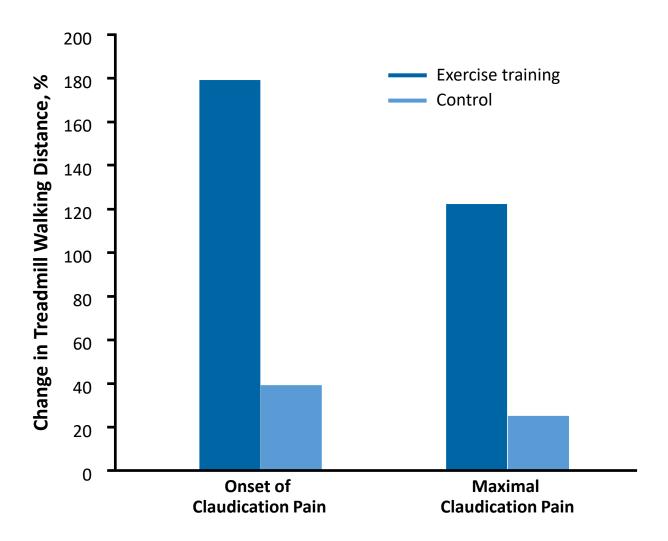
#### Recommendation 15

Pharmacotherapy for symptoms of intermittent claudication

- A 3- to 6-month course of cilostazol should be firstline pharmacotherapy for the relief of claudication symptoms, as evidence shows both an improvement in treadmill exercise performance and in quality of life [A].
- Naftidrofuryl can also be considered for treatment of claudication symptoms [A].



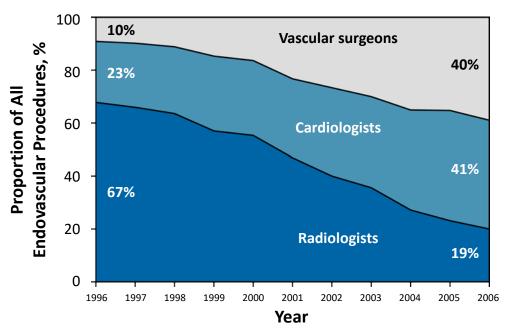
### Effects of Exercise on Claudication: Meta-analysis of 21 Studies





# Revascularization Options

## Who Is Doing Peripheral Interventions These Days?



#### 1.Interventional Cardiologists (41 percent)

- 2. Vascular surgeons (40 percent)
- 3.Interventional radiologists (19 percent)



# PAD: Surgical Revascularization Has Risks

#### Complications

Mortality: 2–5 percent

• MI: 1.9–3.4 percent

Hemorrhage: 2 percent

• Graft thrombosis: 2–7 percent

Wound infection: 8–19 percent

Surgical revision: >20 percent





# Aortoiliac (Suprainguinal) Revascularization

- Initial clinical success of PTA for iliac stenosis exceeds 90%.
- Approaches 100% success rate with focal iliac lesions.
- Technical success of 80-85% for recanalization of long segment iliac occlusions.
- 5 year patency 70-79%
- Factors negatively affecting long term patency
  - Quality of distal runoffs
  - Severity of ischemia
  - Length of diseased segment
  - Female gender



# CFA/SFA Occlusive Disease

- Claudication of thigh, calf or both.
- Calf claudication
  - Most common complaint
  - Cramping pain consistently reproduced with exercise and relieved promptly with rest
  - Differential Diagnosis
    - Nocturnal leg cramps (elderly/nocturnal/rest)
    - Calf pressure and tightness (athletes/chronic exercise)



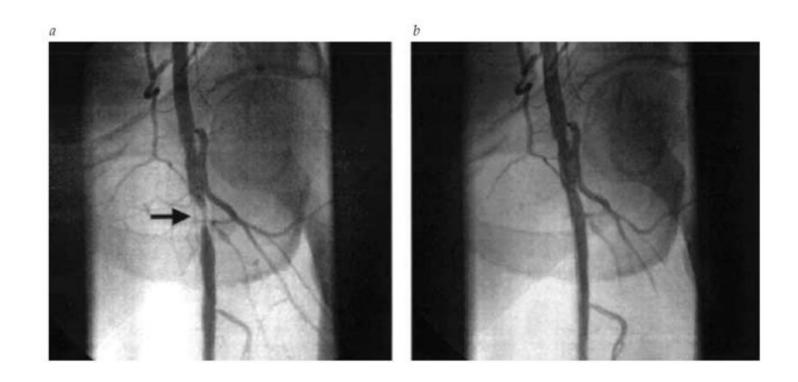








Figure – Right superficial femoral artery is shown here before (left) and after (right) percutaneous transluminal angioplasty.

# Infrainguinal Revascularization

- Endovascular preferred choice of therapy.
- Clinical success rate of PTA 95%
- Predictors of restenosis
  - Length of lesion
  - Extent of outflow disease
  - Tobacco abuse

Table F6. Pooled results of femoral popliteal dilatations

	1-year % patency (range)	3-year % patency (range)	5-year % patency (range)
PTA: stenosis PTA: occlusion PTA+stent: stenosis PTA+stent: occlusion	77 (78–80) 65 (55–71) 75 (73–79) 73 (69–75)	61 (55–68) 48 (40–55) 66 (64–70) 64 (59–67)	55 (52–62) 42 (33–51)

PTA - Percutaneous Transluminal Angioplasty.

# Tibial/Peroneal Occlusive Disease









### Infrapopliteal Endovascular Revascularization

- Indicated for limb salvage.
- Increasing evidence to support a recommendation of PTA in patients with CLI and Infrapopliteal artery occlusion.
- Controversy remains for PTA versus PTA with stent placement.
- Atherectomy starting to play a significant role.



# Thank You



## Need for Clinical Integration



# Why Now – Patient Population Factors

- Aging population.
- Increased prevalence of Diabetics.
- Smoking remains a true healthcare nightmare.
- High percentage of "Asymptomatic" presentation.



## Why now — Economic Factors

- Cost of Amputation.
- Disability claims.
- Cost of prosthesis.
- High mortality and morbidity associated with amputation.



## Why now — Technical Factors

- Tremendous advancements have been made in the medical device industry allowing for better outcomes and limb salvage.
- A few providers from various disciplines have appropriate training and passion to care for these patients.
- A better understanding of "multidisciplinary" approach to treat these patients.
- All team members are of equal significance/importance to save a limb.



#### Where do we start

- Identifying the patients at risk.
- Provider and community awareness to this disease.
- Enhancing risk factor modification.
- Enrolling these patients in surveillance programs.
- Enrolling these patients in PAD rehabilitation programs.



#### Then what

- Identifying the patients at risk.
  - DM/Age/Tobacco abuse
- Enhancing risk factor modification.
  - Consult visit with PAD specialist.
- Enrolling these patients in surveillance programs.
  - Annual Arterial ABI/Dupplex Ultrasound.



# Then what – Asymptomatic Patient

- If patient asymptomatic;
  - PAD Consult.
  - Arterial venous ultrasound.
- Surveillance ultrasounds.
- Monitoring of progression of disease state.
- PAD Rehabilitation/education.
- Supervised exercise walking programs.



# Then what – Symptomatic Patient

- If patient symptomatic;
  - CTA/Angiogram.
  - Interventional procedure.
  - Surgical Bypass.
- Surveillance ultrasounds.
- PAD rehabilitation enrollment.
- Supervised exercise walking programs.

