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Acute Blue Toes: Diagnosis and Plan

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Disclosures

- BD consultant and speaker
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Outline

- What is Blue Toe Syndrome
- What isn't Blue Toe Syndrome (Ddx for blue feet)
- Pathophysiology
- Presentation
- Diagnosis
- Treatment





Blue Toe Syndrome

BTS is:

the acute onset of purple painful digits in the absence of evident trauma, coldassociated injury or disorders that induce generalized cyanosis.





Differential diagnosis of blue toe syndrome

Emboli from cardiac and arterial system	Hypercoagulability disorders	Peripheral vascular pathology and Other
Atrial fibrillation	Antiphospholipid syndrome	Raynaud's phenomenon
Endocarditis	Malignancy	Perniosis (Chilblains)
Left ventricular aneurysm	Thrombocytopenic purpura	Frostbite
Cholesterol crystal from PAD	Disseminated intravascular	Infectious and noninfectious
Penetrating ulcers or Aneurysms of	coagulation	inflammation
the aorto-iliac-femoral arterial system	Warfarin skin necrosis	Medication-induced vasoconstriction
Cardiac/vascular tumors (myxoma)	Cryoglobulinemia	Trauma
Valvular heart disease	Myeloproliferative disorders	Pressure necrosis
Left atrial septal pouch		
latrogenic iniury		

Adapted from: Pradhan et al. Thrombosis Journal (2020) 18:13 https://doi.org/10.1186/s12959-020-00226-x





Perspective and History is Important







Pernio (chilblains)

- Inflamed swollen patches and blistering on the hands and feet
- Primary affects young, underweight women living in cold, humid climates
- Self limiting
- Avoiding cold temperatures and vasospastic factors



STAGES OF CHRONIC VENOUS INSUFFICIENCY







Frostbite







- Painless disorder of decreased oxygen delivery
- Risk factors of acrocyanosis are:
 - cold climate
 - outdoor occupation
 - low body mass index (BMI)
 - major neurological deficit



Acrocyanosis







Raynaud's Phenomenon

- Vasospastic disorder, primarily of the distal extremities
- Attacks can be precipitated by environmental factors, such as exposure to cold or physical trauma. Smoking, heavy alcohol consumption, caffeine, or vasoactive medications can precipitate episodes



The Peripheral Vascular Lab:





Bluish purple ischemic discoloration of the second and fourth toes of the left foot consistent with blue toe syndrome









Epidemiology of BTS

- Process of emboli from proximal lesions produce ischemia in distal arterial beds
- Incidence
 - 0.18% 2.4% in unselected autopsy series
 - 12%-77% in pt's after aortic manipulation
- The risk of atheromatous embolization is directly related to the severity of aortic atherosclerosis



Pathophysiology of BTS

The following 6 key elements are required for the development of cholesterol embolization syndrome:

- 1. Presence of a plaque in a proximal, large-caliber artery (such as the internal carotid artery, the iliac arteries, or the aorta)
- 2. Plaque rupture (spontaneous, traumatic, or iatrogenic)
- 3. Embolization of plaque debris (containing cholesterol crystals, platelets, fibrin, and calcified detritus)
- 4. Lodging of the emboli in small to medium arteries with a diameter of 100 to 200 microns, leading to mechanical occlusion
- 5. Foreign-body inflammatory response to cholesterol emboli
- 6. End-organ damage due to a combined effect of mechanical plugging and inflammation



Clinical Manifestations of Atheromatous Embolization

SKIN

- Purple or blue toes
- Gangrenous digits
- Livedo reticularis
- Nodules

RENAL

- Uncontrolled hypertension
- Renal failure

NEUROLOGIC

- Transient ischemic attack
- Amaurosis fugax
- Stroke
- Hollenhorst plaque



CARDIAC

• Myocardial infarction or ischemia

GASTROINTESTINAL

- Abdominal pain
- Gastrointestinal bleeding
- Ischemic bowel
- Acute pancreatitis

CONSTITUTIONAL SYMPTOMS

- Fever
- Weight loss
- Malaise
- Anorexia

Causes of Blue Toe Syndrome

- Risk factors
 - Established atherosclerosis
 - PaOD
 - HTN
 - Increased age
 - CAD
- Precipitating factors
 - Trauma
 - Vascular surgery
 - Angiographic or endovascular procedures
 - Now the most frequent cause
 - Anticoagulation although controversial
 - Thrombolysis



Atheroemboli

 The thoracic aorta is the source of embolism in 10% to 15% of patients and is associated with nearly a 60% recurrence rate and significant mortality if untreated because visceral ischemia and renal failure are common.





Gross Pathology

(a) Normal thoracic aorta in a child
(b) Severe atherosclerosis of the aorta in an old man.

Thiene G, Basso C, Della Barbera M. Pathology of the Aorta and Aorta as Homograft. *Journal of Cardiovascular Development and Disease*. 2021; 8(7):76. https://doi.org/10.3390/jcdd8070076

(a)

Aortic plaque in the descending thoracic aorta visualized by 2-dimensional TEE



Simple atherosclerotic plaque, measuring 2 mm in this patient.

Complex atherosclerotic plaque with deep ulcerations (arrow).

Complex atherosclerotic plaque with a mobile component (arrow) that represents a thrombus.

Itzhak Kronzon, Muhamed Saric "Cholesterol Embolization Syndrome" 2010 Circulation 631-641, 122, 6 doi:10.1161/CIRCULATIONAHA.109.886465





Findings

- Constitutional signs and symptoms (manifestation of the inflammatory response) such as:
 - Fever
 - weight loss
 - Anorexia
 - Fatigue
 - myalgias are frequent manifestation of the inflammatory response
- Laboratory tests may also show an abnormality in inflammatory markers such as:
 - a rise in leukocyte count
 - erythrocyte sedimentation rate
 - C-reactive protein
 - decrease in serum complement levels (hypocomplementemia)
 - Possible anemia or thrombocytopenia.

Diagnosis

- Histopathological confirmation by biopsy is the only definitive test for cholesterol embolization syndrome
 - Biopsy is performed rather infrequently because it may lead to poor healing at the sampling site
- Clinical Diagnosis
- Peripheral Vascular Lab (PVL)
 - Can present with decreased toe pressures



The Peripheral Vascular Lab: Toe pressures













Asymmetrical flat toe



pressures

Diagnostic imaging

- Unilateral vs Bilateral
 - Bilateral focus above aortic bifurcation
 - CTA C/A/P, Cardiac echo?
 - Unilateral focus below the aortic bifurcation
 - Arterial duplex, CTA AA and runoff, less common angiogram





Characteristic superficial femoral artery atheroembolic lesion responsible for 'blue toe syndrome."



Surgical management of atheroembolization J VASC SURG 1995;21:773-81.

Treatment

- Treatment goals are 2-fold:
 - supportive care for end-organ damage
 - secondary prophylaxis against another episode of cholesterol embolization syndrome.
- Modification of traditional risk factors:
 - Smoking
 - Hypertension
 - serum cholesterol
- Statins weak evidence, but frequently used
- Antiplatelet agents –used presumptively
- ACE inhibitors or direct angiotensin receptor blockers can also be considered.
- Anticoagulation very controversial

BTS presentations



Fig 1. A, Photograph of a foot on presentation, with toe gangrene as manifestation of atheromatous embolization in a patient with AAA and palpable pedal pulses. B, Following stent-graft exclusion of AAA and amputation of gangrenous toes with no further evidence of embolization.



Star-shaped thrombus in small (3.7 cm) abdominal aortic aneurysm that presented with embolization. Characteristic operative findings in aneurysms that embolized was irregular, complex thrombus overlying ulcerative aortic plaque.



Surgical management of atheroembolization J VASC SURG 1995;21:773-81.



a.Toe gangrene as presenting symptom.

b, Chronic organized thrombus retrieved from iliac arteries.

c, Intravascular ultrasound (IVUS) image of eccentric thrombus in the descending thoracic aorta.

d, Preoperative computed tomography

(CT) shows descending thoracic aortic thrombus.

e, Postoperative CT with stent graft coverage of embolizing lesion.

Endovascular strategies for treatment of embolizing thoracoabdominal aortic lesions J Vasc Surg 2014;59:1256-64.

Pradhan et al. Thrombosis Journal (2020) 18:13 https://doi.org/10.1186/s12959-020-00226-x

Intra-cardiac thrombus



Transesophageal echocardiogram (TEE) showing Left atrial septal pouch opening into the left atrial (LA) cavity and a 7.7 × 3mm thrombus (arrow), with the site of attachment corresponding to the LASP. AV, aortic valve; LA, left atrium; RA, right atrium





Presentation

 70 y.o. male with HTN, HLD, ESRD and hx TIA who has been suffering from bilateral lower extremity blue toe syndrome s/p cardiac catheterization for an MI in 5/25. He has progressively declined over the past few months with a 60 lb weight loss in the last 3 months due to loss of appetite. He also states he has no energy and is constantly sleeping because the pain in his feet are so severe. Pt reports some numbness/tingling of his bilateral toes but currently denies any other symptoms.



Physical

- BP 127/76 | Pulse 87 | Temp 95.4 °F (35.2 °C) | Resp 18 | Ht 5' 11" (1.803 m) | Wt 66.8 kg (147 lb 4.3 oz) | BMI 20.54 kg/m2 | SpO2 93%
- General: NAD, AAOx3, thin, Cachectic appearing Caucasian male
- HEENT: NC/AT, PERRLA
- Chest: no chest wall tenderness or deformities
- Lungs: CTAB
- CV: RRR, no M/R/G, no peripheral edema
- Neuro: AAOx3, sensation and motor intact
- Abd: soft, ND, NTD
- Rectal: deferred
- Ext: warm, intact distal pulses, doppler signals present bilaterally. Bluish discoloration on plantar surfaces of bilateral toes with necrotic tissues on plantar surface of right 1st-3rd toes and left 2nd toes

Pulses

Right:

- Carotid nml pulse
- Brachial 2+
- Radial 2+
- Femoral 2+
- Popliteal 1
- Dorsalis 1
- Post Tib 1

Left:

- Carotid nml pulse
- Brachial 2+
- Radial 2+
- Femoral 2+
- Popliteal 1
- Dorsalis 1
- Post Tib 1





Shaggy Aorta









Conclusion

- Many etiologies have similar presentations to true cholesterol embolization syndrome
- Diagnostic studies is based on presumptive cause
- Treatment focuses on supportive care, medical stabilization of plaque and, possibly, treatment of source of embolization