Subclavian Steal Syndrome: An Underrecognized Manifestation of Atherosclerosis - Review of the Current Literature with a Case Report

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Abstract
Subclavian steal is a term used to describe hemodynamics of blood flow in the branches of subclavian artery in patients with an occlusive or stenotic lesion in the proximal subclavian artery. This leads to a retrograde or reversed flow in the ipsilateral vertebral artery supplied by the contralateral vascular bed due to minimal anterograde pressure in the ipsilateral branch. The occlusive lesion is most commonly due to significant atherosclerotic disease. Most patients remain asymptomatic despite having an occlusive lesion. Subclavian steal syndrome (SSS) is the term used when patients develop symptoms related to poor blood supply in the regions supplied by the vertebral artery (posterior circulation symptoms like transient ischemic attacks, vertigo, and dizziness) and/or the brachial artery (arm claudication). Diagnosis is usually made using
Doppler ultrasonography (USG), but other cross-sectional imaging modalities have shown increased prevalence than once thought. Most patients are managed conservatively; however symptomatic patients may need additional endovascular or surgical intervention. Herein, we present a patient of SSS who received subclavian artery stenting followed by complete resolution of symptoms and also review the clinical and management aspects of SSS in detail.

Keywords: Subclavian Steal Syndrome; Atherosclerosis; Stents; Angiography

1. Introduction
Subclavian steal syndrome (SSS) is the phenomenon of reversal of blood flow in a branch of the subclavian artery due to a proximal steno-occlusive lesion in the main vessel [1]. Studies have shown the prevalence of SSS is 0.6-6.4% [2, 3]. With the advancements in the non-invasive diagnostic modalities like Doppler ultrasonography (USG), CT angiography (CTA), and MR angiography (MRA), more asymptomatic cases are being diagnosed to have SSS. The majority of patients are asymptomatic [4, 5]. The steno-occlusive lesion in the proximal part of the subclavian artery leads to impedance in the distal blood flow to its branches namely the vertebral artery, the internal mammary artery, and the axillary artery. With a decrease in the pressure of the blood flow, the vertebral artery can receive blood from the contralateral vertebral artery through the basilar artery or from the carotid artery due to increased pressures in those vessels leading to a reversal in the blood flow in the ipsilateral vertebral artery. This can further be accentuated by exercising the ipsilateral arm which increases the blood flow to the exercising arm further decreasing the blood flow to the vertebral artery [6].

2. Case Study
A 71-year-old female presented to the emergency department with noticeable dizziness, vertigo and pain in her left arm, and left hand for the past 9 months. She also reported her left hand getting cold and shaky sometimes. She had no history of stroke or transient ischemic attack. No significant cardiac history. She had multiple emergency visits in the past due to these symptoms. CT angiography of the head and neck was performed which showed no evidence of intracranial hemorrhage, mass, acute infarct, or intracranial anomaly, however, there was severe stenosis at the origin of left subclavian artery. Further workup with carotid doppler ultrasound revealed new flow reversal in the left vertebral artery, suggesting proximal subclavian stenosis and subclavian steal. Due to her lifestyle being severely impaired by these symptoms, multiple visits to the emergency department for dizziness and posterior circulation issues and having suffered left arm/hand ischemic symptoms she underwent a subclavian angiogram. Angiogram showed patent right subclavian and left common carotid arteries with minimal flow in the left subclavian artery with delayed filling of the subclavian artery through the left vertebral artery suggesting subclavian steal. The left subclavian artery was catheterized with a 5 French Headhunter catheter and 0.035-in angled Glidewire. A baseline left upper extremity angiogram was performed which showed patent left axillary and brachial arteries with slow flow in the forearm and patent radial and ulnar arteries. Measurements were performed from the subclavian artery origin to the takeoff of the left vertebral artery. Subsequently, a 7mm x 37mm Visiproballoon expandable...
covered stent was deployed in the left subclavian artery. Post stenting angiogram performed through the sheath showed brisk flow through the left subclavian artery and antegrade filling of the left vertebral artery. Completion angiogram of the left arm and forearm showed improved flow in left upper extremity. The patient was discharged on clopedogril for 3 months. One-month follow-up imaging showed improved symptoms and excellent radiographic results.

2.1 Background

SSS can present with a spectrum of symptoms depending on the circulation involved but is most commonly diagnosed incidentally as most patients remain asymptomatic. A large clinical study that involved 7881 patients over a period of 6 years undergoing a Doppler ultrasonography (USG) of the cervical region showed the presence of SSS in only 514 (6.5%) patients, out of which only 38 (7.4%) patients were symptomatic [3]. Another study demonstrated that only 5.3% of the 168 patients having SSS under study were symptomatic [7]. A majority (>90%) of patients can be found to have a complete or at least a partial reversal of flow once the caliber of the subclavian artery is reduced by 50%, however, only a small fraction of those patients are symptomatic [8]. Depending on the severity, SSS can be divided into grade I (decreased anterograde flow), grade II (intermittent retrograde flow), or grade III (persistent retrograde flow). Surprisingly, grade II patients are more commonly symptomatic than grade III patients [9].

2.2 Clinical presentation

Upper limb symptoms can include exercise-induced claudication, muscle weakness, paraesthesias, limb coolness, and rest pain in severe cases. The involvement of vertebrobasilar circulation can lead to syncope, ataxia, diplopia, vertigo, and other posterior circulation symptoms [4, 9, 10]. Patients with internal mammary bypass graft can show a reversal of blood flow from the coronary circulation to the internal mammary artery while exercising the ipsilateral upper limb due to increased requirement of blood in the arm and can present with unstable angina [11]. Reversible cognitive deficits in patients with bilateral SSS have been reported secondary to chronic cerebral hypoperfusion [12]. The physical examination can often elicit a weak radial pulse in the affected limb and patients also show a difference in the blood pressures (BP) in the two arms. The BP differences in the two arms have been directly related to the incidence and severity of the symptoms and the need for intervention [10]. Studies have suggested additional workup for SSS in patients with a BP difference of >15mmHg in the two arms. it has been shown that average difference of 36.9 mmHg in the BP of the two arms in patients with moderate stenosis of the subclavian vessel (>50%) [13]. Auscultation may also reveal a subclavian or a suboccipitalbruit due to the turbulent flow [14]. The clinical symptoms can involve the ipsilateral arm, vertebrobasilar circulation, or coronary circulation (in patients with internal mammary artery bypass graft) and occurs due to ischemia due to poor perfusion.

2.3 Diagnosis

2.3.1 Doppler USG: SSS can be diagnosed on imaging with color Doppler USG, CTA, and MRA. Doppler USG of the cervical region is widely recognized as the initial screening test and can demonstrate the blood flow characteristics in the vertebral artery, show reversal of blood flow along with decreased and turbulent flow in the
affected arm, and accentuated peak flow velocity distal to the occlusive lesion in the subclavian artery [5, 15, 16] (Figure 1). In patients with high clinical suspicion showing an absence of retrograde flow in the vertebral artery, a BP cuff test (hyperemia-ischemia test) can be easily performed at the bedside under sonographic examination to reveal any subclinical occult steal at the early stages. In this test, the BP cuff is inflated around the affected arm at a pressure > 20 mmHg than the systolic pressure to cut off the blood supply to the arm. As the cuff is deflated and the blood is preferentially directed towards the ischemic arm, decreased flow or blood flow reversal can be observed on the USG in the vertebral artery in patients with SSS [17]. Kliewer et al proposed that 4 different types of waveforms could be seen based on the degree of steal present. These were classified according to the velocity at the nadir of mid-systolic notch (Figure 2).

![Figure 1](image_url)

**Figure 1:** Left carotid doppler ultrasound in 79-year-old female showing antegrade flow in the left common (a) and internal carotid (b) artery and retrograde flow in left vertebral (c) and subclavian (d) arteries.
Figure 2: Vertebral artery waveforms depending on the degree of steal: Type 1 demonstrating the mid-systolic velocity (MSV) greater than the end diastolic velocity (EDV). Type 2 demonstrating Mid systolic velocity equal to End diastolic velocity. Type 3 demonstrating Mid systolic velocity equal to End diastolic velocity. Type 4 demonstrating retrograde/inverted Mid systolic velocity.

It was greater than the end-diastolic velocity in type 1 waveform, equal in type 2, at the baseline in type 3, and below the baseline in type 4 [17]. However, there are certain limitations of using Doppler USG. The Sonographic examination may not provide a complete visualization of the subclavian and vertebral arteries. Moreover, the calcification in the atherosclerotic plaque might obstruct the view due to the shadowing effect [18]. Studies have shown that USG cannot accurately differentiate between a severely stenosed vessel from a completely obstructed one [19].

2.4 Cross-sectional imaging

The diagnosis is confirmed on CTA and MRA as these modalities help in better characterization, grading and mapping of the lesion with greater detail of the surrounding soft tissue as well [14]. Additionally, angiographic techniques may also help in the diagnosis of additional subclavian artery pathologies like thrombosis or aneurysm [20]. All patients with uncertain or equivocal findings on USG should undergo CTA or MRA. CTA offers a higher spatial resolution and with advanced processing techniques like maximum-intensity projection and multiplanar reconstruction, the subclavian anatomy and the disease extent can be completely assessed [21, 22] (Figure 3).

Moreover, atherosclerosis in the aortic arch and its other branches can also be assessed (seen as vessel-wall irregularities) along with any intracranial disease (Figure 4). It also provides a detailed visualization of calcifications in the aortic arch and other vessels [23]. However, hemodynamic information cannot be evaluated on CTA. MR imaging can help in evaluating the hemodynamics of the blood flow along with the anatomy of the vascular bed. The blood flow dynamics can be assessed using the phase-contrast technique (to identify the direction of blood flow) and the time-of-flight technique (to identify the anatomy of the vascular blood). Additionally, the time-of-flight technique may also be used without intravenous contrast and thus helps in the diagnosis of SSS especially in patients with renal failure [24, 25]. For the overall assessment, the optimal sequence would include a phase-contrast along with contrast-enhanced MRA. 3-D contrast-enhanced MRA has
an even better signal to noise ratio and provides a higher resolution [26]. Angiography is still considered the gold standard for the diagnosis of SSS and also offers the advantage of being a therapeutic procedure as well (Figure 5). It can also help to elucidate the anatomy and patency of collaterals that are being used in different patients [27].

**Figure 3:** Sequential Coronal (a, b, c), sagittal (c, d) and Maximum Intensity Projection (e) reformatted images of CTA in a 79-year-old female showing severe atherosclerotic narrowing in the left subclavian artery (White arrow).

**Figure 4:** Sequential Coronal (a, b, c) and sagittal (c, d, e) reformatted images of CTA in a 79-year-old female at one month follow-up showing patent left subclavian artery stent (White arrow) with eccentric atherosclerotic plaque pushed towards the posterior wall by the stent.
**Figure 5:** Angiographic images in a 79-year-old female. (a) Aortic angiogram showing opacification of the brachiocephalic artery, left common carotid artery and severe stenosis of the left subclavian artery with faint contrast opacification in the subclavian artery (White arrow) (b) Delayed images on the aortic angiogram showing retrograde flow in the left vertebral artery (black arrow) with delayed opacification of the left subclavian and axillary arteries. (c) Selective catheterization and contrast injection through the sheath at the left subclavian artery ostium shows the area of luminal narrowing and tiny nubbin of contrast reflux in the left vertebral artery (Black arrow). This is used for measurement of the stent. (d) Post stenting left subclavian artery angiogram showing improved luminal diameter of the stenotic segment (White arrow) and antegrade flow in the left vertebral artery (Black arrow).

### 2.5 Management

**2.5.1 Medical:** The presence of SSS is highly suggestive of significant atherosclerotic disease in other vascular territories and is associated with an increased risk of total mortality (with a hazard ratio of 1.4) and cardiovascular associated mortality (with a hazard ratio of 1.57) [28]. It warrants a thorough cardiovascular and cerebrovascular workup to look for any significant occlusive atherosclerotic lesions. Asymptomatic patients and patients with mild symptoms are usually managed medically which include aspirin, β-blockers, ACE-inhibitors, and statins, a regimen that is similarly used to manage atherosclerosis in other vascular beds along with lifestyle modification for hypertension, diabetes mellitus, smoking, and alcohol use.
It has been shown that only 1.4% of patients with SSS require any therapeutic intervention [3].

2.6 Endovascular and surgical

Traditionally treated with a surgical bypass or a transposition technique, the management of SSS has been completely transformed with endovascular treatment introduced by Kim and Bachman in 1980 [30]. Percutaneous transluminal angioplasty (PTA), or balloon angioplasty, can be performed with or without the placement of stents. PTA has proven to be an excellent alternative to open vascular surgery with a success rate of nearly 95% and comparable long-term patency rates (80-90%) with a much lesser incidence of complications that are seen with surgical options. Previous studies have shown that only 10% of the patients develop restenosis during long-term follow-up and nearly 95% of those lesions can be successfully managed with a repeat PTA [31, 32]. Placement of stents along with PTA can further improve the clinical outcomes in patients with SSS [33]. The algorithm “stent first, then surgery” has garnered immense popularity [34]. Angioplasty without stenting (primary or secondary) was the initial modality tested, with Hebrang et al reporting a technical success rate of 93% and a clinical success rate of 86.5% and 78.8% at 1 and 3 years respectively in a cohort of 52 patients [35]. Despite its early success, this procedure had its share of drawbacks namely a less favorable outcome for a total occlusive disease of the subclavian artery and a considerably high rate of restenosis. Duber et al reported a 57% rate of restenosis or reobstruction in treated patients [36]. To combat this problem, angioplasty with stenting was introduced in the 1900s as a reasonable alternative. The proposed advantages of stenting included prevention of elastic recoil of atherosclerotic lesion and reduction in thrombotic or embolic complications by trapping of the exposed intraluminal plaque [30]. Considerable success was achieved with Sakai et al reporting 89.3% primary and 100% secondary patency rates [37].

However, further randomized controlled clinical trials are required to compare angioplasty with or without stenting to draw a definite conclusion. Irrespective of the technique used, a regular follow-up for 2 years post endovascular therapy is recommended [38]. Drug-eluting stents have also been introduced recently to improve the patency rates. One of the major challenges faced while performing the endovascular treatment is the difficulty in traversing the vessel for the management of chronically occluded subclavian artery. Multiple techniques have been tried to solve this problem, namely bi-directional access, extra-stiff catheters, and an iatrogenically created sub-intimal dissection [30]. Some authors have reported techniques like non-compliant and compliant balloon catheters and distal filters to prevent distal emboli, but that discussion is beyond the scope of this manuscript [39, 40]. Another important thing to consider is the disparity between technical and clinical success with regard to endovascular procedures. This could be due to the fact that the diameter change (morphological parameter) does not always correlate with hemodynamic status (pathophysiological parameter). Soga et al reported the technical success rates for SCA occlusion and stenosis as 100% and 91% respectively, but clinical success rates were shown to be 92% and 80% respectively [41]. A technique to assess hemodynamic status during endovascular therapy for SSS was used by a Chinese study group in 11 patients with SSS undergoing angioplasty with or without stenting. They used a pressure wire to detect the trans-stenosis pressure difference during the procedure, a
technique that has been used previously for coronary, renal, and mesenteric vascular beds. They concluded that the pressure difference is closely related to clinical symptoms and degree of steal with values of 6-9 mm of Hg for grade 1 and 2 steals and >10 mm of Hg for ≥grade 3 steal. It was also reported that the gradient of <3 mm of Hg was associated with high post-procedural clinical success [34]. Patients who are not amenable to endovascular treatment require surgical options. The surgical vascular interventions include bypass (axillary-axillary or carotid-subclavian with latter showing lower mortality and recurrence rates) and transposition of SA [38]. Studies have shown that the transposition of SA has the highest long term patency rate (100% at 5 years) [17].

3. Conclusion
SSS is a hemodynamic phenomenon that causes decreased perfusion in the branches of the subclavian artery and retrograde flow of blood in the vertebral artery due to decreased anterograde flow caused by an occlusive lesion in the subclavian artery. An overwhelming majority of patients remain asymptomatic and those with severe occlusion can present with symptoms involving the vertebrobasilar circulation or ipsilateral arm ischemia. A thorough workup for SSS and other atherosclerotic lesions should be done in all such patients. Most patients can be managed conservatively by medical management. Symptomatic patients are managed with endovascular (PTA with or without stenting) first or surgical management in select cases.

References


