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Original Article

Pipeline embolization device as primary treatment for cervical internal carotid artery pseudoaneurysms

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Received: 24 June 16  Accepted: 15 November 16  Published: 19 January 17

Abstract

Background: Limited data exists on the durability and occlusion rate of treating extracranial cervical internal carotid artery pseudoaneurysms using the pipeline embolization device (PED) flow-diverting stent.

Methods: Three patients presenting with dissecting cervical internal carotid artery pseudoaneurysms were treated with the PED as the sole treatment modality.

Results: In all three patients, successful aneurysmal occlusion and parent vessel reconstruction occurred on immediate angiography and continued on 6-month follow-up. No immediate or delayed complications were seen, and all patients remained neurologically intact.

Conclusion: Complete aneurysmal occlusion and long-term angiographic occlusion can occur after PED treatment of cervical carotid pseudoaneurysms. In select patients, the PED can be a suitable primary treatment modality with good neurological outcome for cervical carotid pseudoaneurysms.

Key Words: Cervical carotid, dissection, pipeline embolization device, pseudoaneurysm

INTRODUCTION

Cervical internal carotid artery dissections and pseudoaneurysms are not uncommon. Recent studies have demonstrated a prevalence of blunt cervical vascular injury at 1.2–1.6% of trauma patients and 2.7% of multisystem trauma patients. The average annual incidence of spontaneous and iatrogenic injuries to the carotid and vertebral arteries has been reported at 2.9 per 100,000.

The majority of dissections, Denver Grades 1 and 2, heal with medical therapy and less than 10% progress to a pseudoaneurysm. Grade II lesions improved to grade I lesions in 13.5% of the patients whereas 45.9% progressed to a worse grade. The majority of pseudoaneurysms persist, with only 3% of Grade 3 lesions healing with medical therapy and another 3% progress to complete vessel occlusion. Studies with early follow-up imaging show that over 90% of pseudoaneurysms persisted at 1 week. In a study with 6-month follow up, 72% of grade I lesions and 33% of grade II lesions healed. Edwards et al. reported that, with medical therapy, only 10% of pseudoaneurysms healed whereas 40% progressed.

How to cite this article: Wang A, Santarelli J, Stiefel M. Pipeline embolization device as primary treatment for cervical internal carotid artery pseudoaneurysms. Surg Neurol Int 2017;8.

http://surgicalneurologyint.com/Pipeline-embolization-device-as-primary-treatment-for-cervical-internal-carotid-artery-pseudoaneurysms/
The recent advances in endovascular technology have shifted treatment paradigms away from microsurgical techniques and towards endovascular treatment of dissecting carotid pseudoaneurysms. A few small retrospective case series have demonstrated the success of vascular stents and coils in treating these lesions. Flow-diverting devices such as the pipeline embolization device (PED; ev3-Covidien, Irvine, CA, USA) represent a changing paradigm in the treatment of intracranial aneurysms. While specifically designed for use on wide necked unruptured large and complex aneurysms, the PED has found success in treating internal carotid artery and vertebral artery dissecting pseudoaneurysms. The majority of the current literature supporting PED success has been centered on treating the intracranial segment of these vessels.

The application of the PED on cervical carotid pseudoaneurysms is limited. To date, there has only been one published study applying the PED to treat high cervical carotid pseudoaneurysms. We present our experience using the PED as the sole treatment modality in three patients with high cervical carotid pseudoaneurysms, and briefly discuss the technical and perioperative management of these patients.

**CASE REPORTS**

Three patients with Denver Grade 3, dissecting cervical carotid artery pseudoaneurysms were treated at our institution using the PED [Table 1]. A pretreatment computed tomography (CT) of the head and CT angiogram of the head and neck were performed on all patients. All cases demonstrated a dissecting pseudoaneurysm of the extracranial cervical internal carotid artery segment. The clinical status with respect to presenting symptoms and Glasgow Coma Score (GCS) was recorded prior to endovascular treatment.

All patients were loaded on aspirin and clopidogrel 24 hours prior to PED placement with platelet reactivity testing performed to confirm adequate platelet inhibition. At the beginning of each case, a baseline activated clotting time (ACT) was obtained, and a weight based intraoperative bolus of intravenous heparin was given after femoral artery access to maintain an ACT of >200 seconds throughout the procedure. All interventions were performed in a biplanar fluoroscopic angiography suite under general anesthesia using intraoperative monitoring (somatosensory evoked potentials, motor evoked potentials, and electroencephalography).

The PED (ev3-Covidien) was used in all cases as the only device. Conventional and three-dimensional (3D) angiographic studies were performed and working angle projections for PED placement were acquired at high magnification of the cervical carotid artery. In all patients, consecutive PED stents were placed in a telescoping fashion, distal to proximal, to cover the pseudoaneurysm.

Immediately after the treatment, all patients were extubated and observed in the intensive care unit for 24 hours with hourly neurological examinations. All patients were verified to be aspirin and clopidogrel responders and were continued on daily 325 mg aspirin and 75 mg clopidogrel.

The angiographic features of the pseudoaneurysms and outcomes with respect to clinical and radiographic images of the patients are reviewed. With radiographic outcome, immediate and 6-month follow up angiograms of the patients were reviewed to examine contrast stagnation, occlusion of the pseudoaneurysm, and in-stent patency following PED deployment.

**Case 1**

Patient 1, in the second decade of life, presented intubated with multisystem trauma from a motor vehicle accident; GCS 5T and an injury severity score (ISS) of 41/75. Initial CT angiogram of the head and neck demonstrated grade I mild focal narrowing of the right cervical carotid artery at the level of C1-2. A repeat CT angiogram 1 week later showed stability of the diseased carotid segment whereas a CT angiogram at 3 weeks demonstrated interval development of a 6.5 mm × 14 mm dissecting pseudoaneurysm of the right cervical internal carotid artery at the C1-2 level [Figure 1a]. Cerebral angiography was performed [Figure 1b and c] and the aneurysm was treated with 4 telescoping PEDs (two 5.0 mm × 20 mm, 5.0 × 18 mm, 4.25 mm × 20 mm). Final angiograms showed good wall apposition of the PEDs and improved vessel flow and caliber [Figure 1d and e].

**Table 1: Demographic and clinical characteristics, treatments, and outcomes in three patients**

<table>
<thead>
<tr>
<th>Patient</th>
<th>Age (years), sex</th>
<th>Etiology, Presentation</th>
<th>Side</th>
<th>Aneurysm size</th>
<th>Location</th>
<th>Treatment</th>
<th>6-month Follow-up</th>
</tr>
</thead>
<tbody>
<tr>
<td>1</td>
<td>20s, M</td>
<td>Iatrogenic injury, polytrauma</td>
<td>Right</td>
<td>2.0 × 1.0 mm</td>
<td>Distal cervical</td>
<td>Pipeline×4</td>
<td>Complete occlusion</td>
</tr>
<tr>
<td>2</td>
<td>40s, F</td>
<td>Iatrogenic injury, Horner’s syndrome</td>
<td>Right</td>
<td>5.0 × 3.5 mm</td>
<td>Distal cervical</td>
<td>Pipeline×2</td>
<td>Complete occlusion</td>
</tr>
<tr>
<td>3</td>
<td>20s, M</td>
<td>Iatrogenic injury, polytrauma</td>
<td>Right</td>
<td>15.0 × 10 mm</td>
<td>Distal cervical</td>
<td>Pipeline×4</td>
<td>Complete occlusion</td>
</tr>
</tbody>
</table>
Case 2
Patient 2, in the fourth decade of life, presented with clinical symptoms consistent with recurrent Horner’s syndrome after recovering from an upper respiratory infection. CT angiogram of the neck demonstrated an enlarging medially projecting 3.3 mm × 6.6 mm pseudoaneurysm of the right cervical internal carotid artery compared to the CT angiogram 3 years ago. Cerebral angiography confirmed the pseudoaneurysm. Two telescoping PEDs, 4.5 mm × 16 mm and a 5.0 mm × 16 mm, were used.

Case 3
Patient 3, in the second decade of life, presented intubated with multisystem trauma from a motor vehicle accident; GCS 6T with extensive extremity fractures and lung injuries with an ISS of 48/75. Initial CT angiogram of the head and neck demonstrated a dissection flap with associated pseudoaneurysm of the right distal cervical internal carotid artery. Follow up CTA showed that the pseudoaneurysm enlarged. Four telescoping PEDs, 5.0 mm × 20 mm, were used.

PEDs were successfully placed in all three patients. In patients 1 and 3, overlapping PEDs were deployed. In patient 2, a second PED was needed after foreshortening of the first PED was observed. At the completion of the cases, increased intra-aneurysmal contrast stagnation, improved vessel caliber, and normal intracranial vascular flow were seen on the final angiographic runs.

No neurological deficits or access site complications were observed. Patients 1 and 3 underwent multiple surgeries to repair severe facial and lower extremity fractures and remained neurologically intact at the time of discharge. Patient 2 continued to exhibit Horner’s syndrome at the time of discharge. All patients were discharged on daily aspirin and clopidogrel therapy.

Patient 1 returned within 1 week of discharge with transient left-sided weakness. A CT head confirmed no delayed intracerebral hemorrhage, and a CT angiogram demonstrated patent intracranial vessels without in-stent thrombosis. Magnetic resonance imaging of the brain showed no restricted diffusion changes. The patient was observed and discharged after complete resolution of his symptoms.

All patients remained neurologically intact (modified rankin scale = 0) at 6-month follow up visit. Complete aneurysm occlusion and normal vessel caliber were seen on repeat angiography at the time.

DISCUSSION
The majority of extracranial dissections, Denver grade 1 and 2, heal with medical therapy alone, however, medical therapy is not as effective for pseudoaneurysms, Denver Grade 3. Endovascular treatment of Grades 1 and 2 dissections is usually reserved for patients who fail medical therapy or who have symptoms from perfusion abnormalities. Endovascular treatment of pseudoaneurysms is recommended for those with enlarging pseudoaneurysms or those with embolic events despite medical therapy. Endovascular constructs offer several advantages, they restore the vessel lumen of the dissected segment, and provide a support scaffold for endothelialization and healing. Moreover, depending on the porosity of the stent or construct, it reduces or
Several case reports have described the Balloon-expandable stent's (Boston Scientific Corporation, Fremont, CA, USA) thin and open cell frame design leads to a relatively low metal surface area and may be suboptimal in covering the porous inflow zone to a pseudoaneurysm and being used as reliable stand-alone therapy. However, in two studies, the Neuroform stent was used with success in distal cervical dissections.[12,17] Coldwell et al. described 14 traumatic carotid artery pseudoaneurysms treated with self-expanding bare metal stents with all aneurysms remaining occluded at 6 months.[13] Recently, Ansari et al. used a Neuroform self-expanding stent to treat nine carotid pseudoaneurysms with complete or near-complete occlusion of the aneurysms.[2]

**Current applications of the pipeline embolization device for cervical and intracranial dissections**

The PED is a microcatheter delivered self-expanding stent-like construct that is very effective for treating intracranial fusiform, large, or wide-necked aneurysms. The PED offers the advantage of being used as a stand-alone device with immediate flow diversion and intra-aneurysmal thrombosis.

The application of the PED on treating dissecting pseudoaneurysms has mainly focused on the intracranial circulation.[11,14,21,27,35] De Barros Faria et al. demonstrated the successful application of the PED alone on 21 dissecting pseudoaneurysms of the posterior circulation, in which 87.5% were occluded at 3-month follow-up.[15] Similarly, Narata et al. and Yeung et al. both reported complete occlusion of all ruptured dissecting vertebral aneurysms treated in their respective case series with the PED stent alone.[24,37] A few case reports have been reported in the literature highlighting the use of the PED on anterior circulation. Amenta et al. was the first to highlight the successful use of the PED in the treatment of cavernous carotid pseudoaneurysm incurred during transphenoidal pituitary tumor resection.[13] Subsequently, Nerva et al. recently discussed their success with the PED in treating two iatrogenic carotid pseudoaneurysms of the cavernous segment.[36]

The PED has several properties that make it suitable for treating extracranial cervical dissections. Its flexible self-expanding design combined with the advantage of using a smaller guide and microcatheter platform allows for greater maneuverability through tortuous and narrowed vessel segments at the skull base. In their series of 11 patients with 13 extracranial carotid dissections, Brzezicki et al. demonstrated immediate complete revascularization in 91% of the vessels and complete occlusion of 50% of the associated pseudoaneurysms.[7] Follow-up imaging of nine vessels at 9 months demonstrated good durability without significant in-stent stenosis or migration of the PED construct. Park et al. described one patient who was treated with a PED for an iatrogenic cervical carotid...
dissection, which on follow-up angiogram 2 days later showed complete occlusion.\textsuperscript{[11]} Cohen et al. showed good long-term durability at 18 months in their cohort of three patients treated with the PED for cervical vertebral artery dissections.\textsuperscript{[12]} Fischer et al. described their experience with the PED in treating 12 cervical carotid lesions in their larger cohort of 69 lesions.\textsuperscript{[20]} To date there is little data on the details of occlusion rates or long-term durability with this approach.

**Technical considerations in choosing a stand-alone construct**

Stent reconstruction in the distal cervical carotid artery may be challenging because of the tortuosity of this segment and the changing diameter and mobility of the cervical carotid artery with head movements. There has been limited use of the PED on extracranial cervical carotid pseudoaneurysms. In our three cases, conventional self-expanding carotid or smaller biliary stents would have necessitated larger constructs being navigated through the dissected segment and into the petrous carotid segment. The PED’s flexible and self-expanding design combined with the advantage of using a smaller guide and microcatheter platform allows for greater maneuverability through tortuous and narrowed vessel segments.\textsuperscript{[25,29,30,33]}

The length and diameter of the diseased segment as well as the adjacent normal segments are carefully evaluated when choosing the appropriate stent or flow diverter construct. Stents such as the Neuroform and Enterprise (Codman Neurovascular, Raynham, MA, USA) have a maximum diameter of 4.5 mm, making them less suitable in the larger mid-cervical and proximal-cervical internal carotid artery. The PED is available from 2.5 to 5.0 mm with 0.25 mm increments and lengths of 10 to 35 mm. The 5.0 mm PED expands to 5.25 mm.\textsuperscript{[29]}

It is important to consider that oversizing the PED can lead to elongation of the device with poor coverage of the lesion and a decrease in flow diversion. Undersizing the device may result in poor wall apposition, an endoleak, as well as a potential risk of stent migration. Selection of a suboptimal PED and potential stent foreshortening and delayed migration of the device in both intracranial and extracranial disease has been reported.\textsuperscript{[9,10,13,23]} Rahal et al. used the PED in conjunction with an anchoring Solitaire stent to treat two patients with shallow wide-necked cervical pseudoaneurysms in the hopes of avoiding stent migration and foreshortening.\textsuperscript{[14]} They deployed a Solitaire stent (ev3 – Cavidien, Irvine, CA, USA) in a concentric manner to anchor their PEDs, but still experienced delayed foreshortening of the PED on follow-up imaging. In all three of our patients, we placed multiple stents in tandem and ensured that the stent constructs were long enough to guarantee coverage of the entire lesion with sufficient overlap of the normal proximal and distal adjacent segments. Six-month follow-up angiograms showed intact coverage of the aneurysm neck, good wall apposition of the PED, and no evidence of aneurysm recurrence in all three patients.

The effectiveness of flow diversion may vary with the amount of metal coverage. The high porosity of currently available stents limits the ability to change inflow within an aneurysm. The PED was specifically designed to address this issue with the goal of promoting more effective flow diversion. The PED achieves this because of its low porosity and 30–35% surface metal area coverage, which is more than other stents such as the Xpert Biliary Stent (Abbott Vascular, Santa Clara, CA, USA) (20%) and Neuroform (9%) and Enterprise Stents (7%) [Table 2]. As a comparison, this results in a flow reduction of 37% and 55% for single and overlapping PED vs. a flow reduction of 24% in the Neuroform Stent.\textsuperscript{[6,17]}

Lastly, while the PED offers certain advantages over other stents available in the market, it is associated with adverse cost considerations ($12500 per PED) and may not be ideal for each patient and each institution, especially if multiple PEDs will be placed.

**CONCLUSIONS**

The application of flow diverters in treating cervical carotid pseudoaneurysms is limited. Our study highlights the use of the PED as the sole modality for the treatment of cervical carotid artery dissecting pseudoaneurysms. Although small, our series demonstrates the successful use of the PED as a primary endovascular modality with excellent 6-month radiographic and clinical outcomes.

**Financial support and sponsorship**

Nil.

**Conflicts of interest**

There are no conflicts of interest.

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