

## Review

# Endovascular treatment of the cavernous sinus dural arteriovenous fistula: current status and considerations

Kun Hou<sup>1\*</sup>, Guichen Li<sup>2\*</sup>, Tengfei Luan<sup>1</sup>, Kan Xu<sup>1</sup>, Jinlu Yu<sup>1</sup>✉

1. Department of Neurosurgery, The First Hospital of Jilin University, Changchun, 130021, China

2. Department of Neurology, The First Hospital of Jilin University, Changchun, 130021, China

\*Kun Hou and Guichen Li contributed equally to this manuscript, and they are co-first authors.

✉ Corresponding author: Jinlu Yu, E-mail: jlyu@jlu.edu.cn, Department of Neurosurgery, The First Hospital of Jilin University, 1 Xinmin Avenue, Changchun 130021, China. ORCID: Jinlu Yu, 0000-0003-2329-7946; Kun Hou, 0000-0001-7112-7883

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## Abstract

A cavernous sinus dural arteriovenous fistula (CS-DAVF) is an abnormal arteriovenous communication involving the dura mater within or near the CS wall. The dural arteries from the internal carotid artery and external carotid artery supply the CS-DAVF, and the superior ophthalmic vein (SOV) and inferior petrous sinus (IPS) are frequent venous drainers. In CS-DAVF cases, high-risk lesions require treatment. Endovascular treatment (EVT) has been the first-line option for CS-DAVFs. To our knowledge, a review of the EVT of CS-DAVFs is lacking. Therefore, in this paper, we review the available literature on this issue. In addition, some illustrative cases are also provided to more concisely expound the EVT of CS-DAVFs. According to the recent literature, transvenous embolization via the IPS is considered the most effective method for EVT of CS-DAVFs. In addition, the transorbital approach is another reasonable choice. Other venous approaches can also be tried. Because of the low cure rate, transarterial embolization for CS-DAVFs is limited to only highly selected patients. In the EVT of CS-DAVFs, various agents have been used, including coil, Onyx, and n-butyl cyanoacrylate, with coil being the preferred one. In addition, when EVT cannot obliterate the CS-DAVF, stereotactic radiotherapy may be considered. In general, despite various complications, EVT is a feasible and effective method to manage CS-DAVFs by way of various access routes and can yield a good prognosis.

Key words: cavernous sinus, dural arteriovenous fistula, endovascular treatment, review

## Introduction

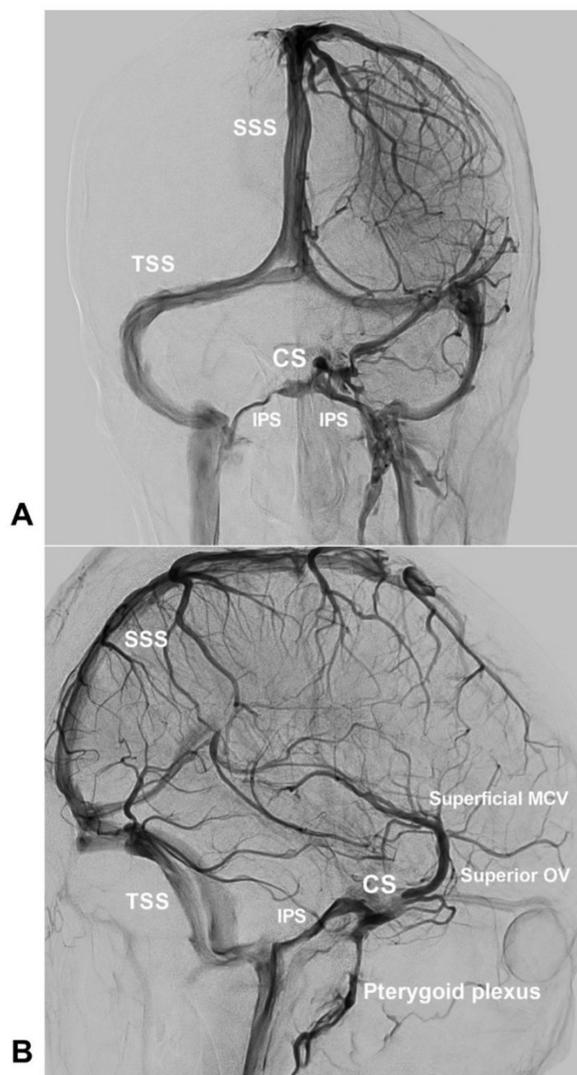
Cavernous sinus dural arteriovenous fistula (CS-DAVF) is the abnormal communication between the CS and the dural arteries from the internal carotid artery (ICA) and external carotid artery (ECA), commonly caused by CS recanalization after thrombosis [1].

In CS-DAVFs, retrograde arterialized flow in the draining veins results in characteristic symptoms [2]. Low-flow asymptomatic CS-DAVFs can close spontaneously, but high-flow symptomatic CS-DAVFs can rarely disappear spontaneously [3]. Therefore, treatment is recommended for symptomatic CS-DAVFs. Currently, endovascular treatment (EVT) has become the first-line choice for CS-DAVFs [4]. In addition, adjuvant radiosurgery may be considered after unsatisfactory EVT [5].

Currently, a comprehensive review of EVT for CS-DAVFs is still lacking. In this paper, we review the available literature on this issue. In addition, some illustrative cases are also provided to more concisely expound the EVT of CS-DAVFs.

## Critical vessels in the CS region

The CS is an extradural space located in the bilateral parasellar compartment. The ICA passes through the CS and gives rise to several small branches supplying the dura. In addition, the ECA also supplies the CS through its dural branches [6, 7]. The blood supply of the CS can be bilateral [8]. There are constant anastomoses between the ICA and ECA, which are called “dangerous anastomoses” [9].



**Figure 1. Normal venous drainage of the CS.** An angiogram of the left ICA of the venous phase in AP (A) and lateral (B) views shows the CS and the venous structures around the CS. **Abbreviations:** AP, anteroposterior; CS, cavernous sinus; ICA, internal carotid artery; IPS, inferior petrous sinus; MCV, middle cerebral vein; OV, ophthalmic vein; SSS, superior sagittal sinus; TSS: transverse sinus.

CS is not a single venous lake but a venous plexus. It receives venous drainage from the superior and inferior ophthalmic veins (OVs) anteriorly and from the sphenoparietal sinus and the superficial middle cerebral vein laterally, and drains posteriorly to the superior and inferior petrous sinuses and basilar plexus, inferiorly to the pterygoid plexus [10, 11]. The bilateral CSs communicate with each other through the intercavernous sinus [8, 12].

The normal venous drainage of CS is shown in Figure 1.

## Angioarchitecture and classification of CS-DAVF

### Feeding artery

Feeders of CS-DAVFs are from abundant ECA and/or ICA branches, in which the middle meningeal

artery (MMA), accessory meningeal artery, and ascending pharyngeal artery (APhA) are the most clinically significant branches [13-16]. These feeders may be bilateral in nearly 70% of cases [16, 17]. Under normal circumstances, these small dural branches may not be visible on catheter angiography but become hypertrophic or enlarged when supplying CS-DAVFs [18].

### Fistula point

The CS has several compartments, but usually, the fistulous portion does not involve the entire CS; the fistulous portion is also called the shunted pouch (SP) [19-22]. The SP is a tubular or elliptical structure that is separated from the main sinus lumen and the site where multiple feeding arteries converge and continue to the CS [23].

The SP is mainly situated at the posterior compartment of the CS and connects with the intercavernous sinus. It might have one or more of the following angiographic findings: a) an early opacified area in the early arterial phase of catheter angiography, b) a jellyfish-like sign denoting the branches of arterial supply converging at the fistula point, c) changes in the density of the contrast medium from dark in the artery to gray in the veins on the fistula point, d) enlarged and tortuous arterial feeders that can directly point to the fistula point [24, 25].

The SP may be single or multiple, and it can be divided into dural, extradural, or osseous types based on the converging site of arterial feeders [17, 26].

### Venous drainage

In the high pressure gradient environment of CS-DAVFs, the arterialized veins reverse the direction of their normal inlet, and the drainers are highly variable [16]. Superior ophthalmic vein (SOV) drainage is a frequent finding that can be present in 88% of patients with CS-DAVFs, while the rate is only 42% with inferior petrous sinus (IPS) drainage and 34% with cortical venous drainage [27].

The contralateral or bilateral drainage can also be visualized [13, 16]. Rarely, venous reflux from CS can drain into the posterior fossa and brainstem by bridging veins [28]. The draining vein can be stenotic or obliterated due to thrombus formation [27, 29-31].

### Classification

The Borden and Cognard grades are most commonly used for the classification of DAVFs. Their grading criteria are based on the venous drainage pattern, which is also applicable to CS-DAVFs [32-34]. However, the CS-DAVF has its own anatomical and angiographic characteristics and should be classified more specifically [35-37]. The following classification

systems are helpful to guide the EVT of CS-DAVFs [38-41].

In 2005, Suh et al. divided the CS-DAVF into 3 types: proliferative CS-DAVF receiving numerous arterial feeders with rapid filling of the CS and drained by multiple venous outlets, late restrictive CS-DAVF draining primarily into the superior and inferior OVs, and restrictive CS-DAVF, which is the transitional type between the proliferative and late restrictive CS-DAVFs [38].

In 2015, Thomas et al. divided CS-DAVFs into 5 types, with types 1-4 designed for DAVFs [39]. Type 1 has posterior/inferior drainage (IPS, superior petrous sinus (SPS), pterygoid plexus). Type 2 has posterior/inferior and anterior drainage (superior and inferior OVs). Type 3 has anterior drainage only. Type 4 has retrograde drainage into cortical veins.

In 2017, Wenderot et al. divided the CS-DAVF into 3 types: type 1 has free IPS outflow from the affected CS; type 2 has bilateral CS drainage but unilateral IPS occlusion, and type 3 has no IPS outflow at all [40].

These classification systems provide useful information on fistula anatomy and hemodynamics, which could further aid in the planning of fistula access during EVT [42].

## Outline of EVT

Currently, various EVTs are available for CS-DAVFs, mainly including transfemoral transvenous embolization (TVE) and transarterial embolization (TAE), a transorbital puncture approach, and direct puncture or surgical exposure to obtain transvenous access to the CS [3, 5, 43].

In CS-DAVFs, TAE is usually difficult because most CS-DAVFs are fed by numerous and tiny dural branches from the ICA and ECA. In addition, TAE has a low cure rate. Therefore, TAE should be used only in highly selected cases [13, 44].

Nevertheless, TVE has become the mainstream EVT for CS-DAVFs, of which the transfemoral IPS approach is the favorite, shortest, and most direct route to the CS [1, 15]. In TVE, precise identification of the SP location is crucial because targeted dense embolization of the SP can reduce the amounts of embolic agents used and thus avoid complications [24, 25, 45].

If the transfemoral IPS approach is inaccessible, an alternative transorbital approach can be tried, which includes direct puncture of the superior and inferior OVs and CS as well as puncture of the superior OV after surgical exposure [5].

In the EVT of CS-DAVFs, various materials have been used, including coil, Onyx (Medtronic, Irvine, California, USA), and n-butyl cyanoacrylate (NBCA)

[46]. For TVE, coil is the preferred material, and Onyx can also be used in combination or alone [3]. For TAE, NBCA is an effective agent due to its good ability in distal penetration [13]. However, Onyx is also a promising choice due to its nonadhesive nature, allowing for a longer injection time [47].

## TVE

### Transfemoral venous approach

For the CS-DAVF with a visible IPS, trans-IPS access was successful in nearly 90% of the cases [30, 48]. The contralateral IPS can also serve as an alternative route [15, 49]. 3D rotational venography of the internal jugular vein (IJV) is very useful for revealing the details of the IPS and creating a plan to pass it [43, 50]. In addition, preoperative CT angiography (CTA) can also help to localize the IPS [51].

To perform TVE, the reopening of an occlusive IPS is feasible, and the success rate ranges from 50% to 90% [15, 16]. The major factors affecting the reopening of an occlusive IPS are the duration and length of occlusion. It is difficult to reopen an IPS with a long duration and length of occlusion [16, 43]. During reopening the IPS, a standard hydrophilic-polymer-jacketed 0.035-inch guidewire can advance as a frontier wire for probing the IPS. Then, the trace of the guidewire on the roadmap image can serve as a guide for microcatheter navigation through the IPS [27, 50]. In addition, the microwire looping technique for breaching the occluded IPS under the reinforcement of guiding system support is also helpful, during which the 5F catheter fixed to the outlet of IPS provides a stable support [29, 51, 52].

In addition to targeted dense SP packing, the following details should also be considered: 1) the origin of the superior and inferior OVs should be packed to ensure that the fistula is not redirected into the orbit, and 2) cortical and brainstem venous drainage should be occluded with priority [27, 29, 30, 38, 53].

If the SP packing is not satisfactory or the CS-DAVF belongs to the proliferative type according to Suh et al.'s classification, the entire CS has to be packed densely [38]. A case of CS-DAVF with entire CS packing via IPS is illustrated in Figure 2.

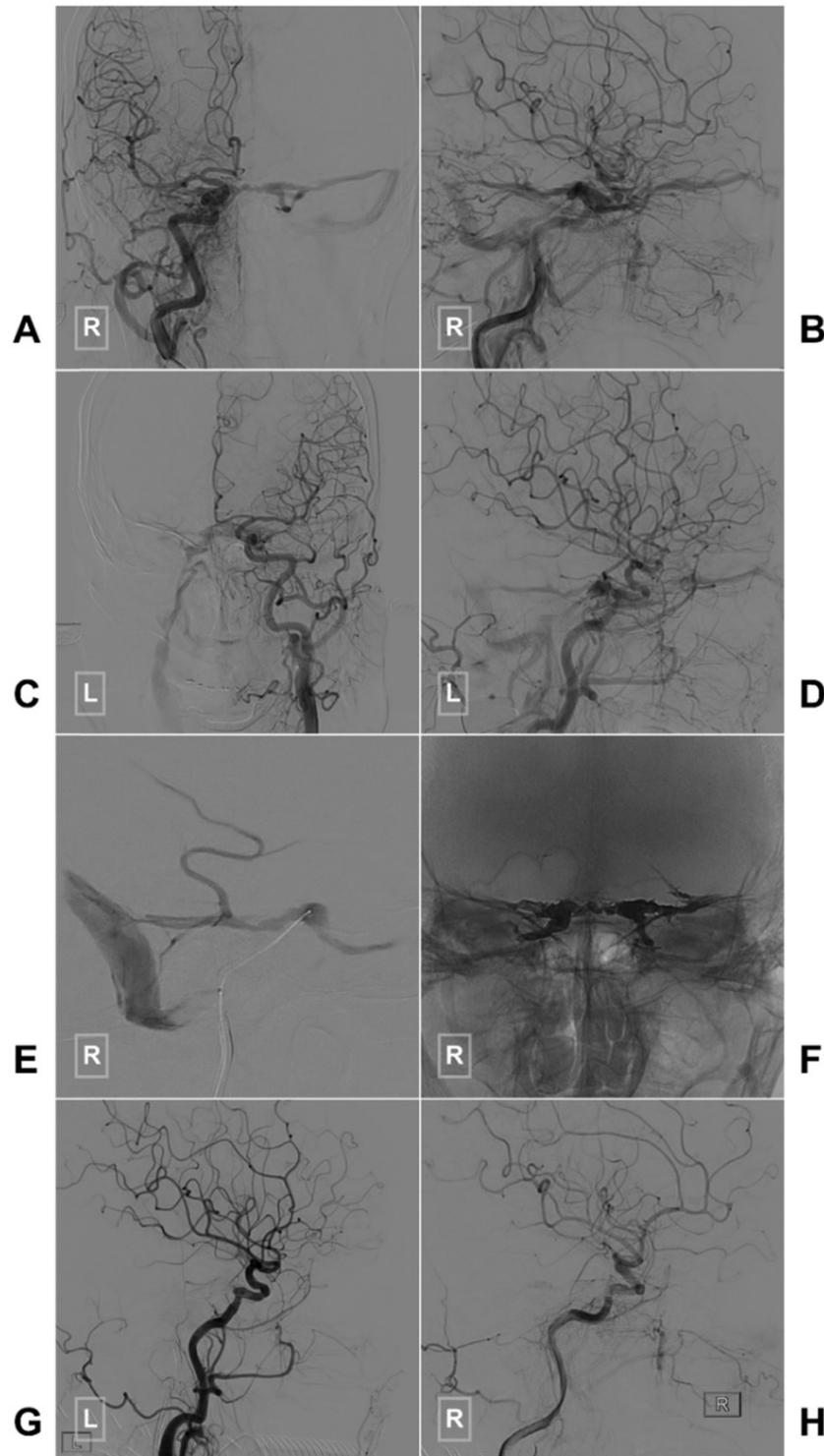
In addition to the trans-IPS approach, other transfemoral routes can also be used to access the CS, e.g., the transpterygoid plexus approach [48, 54-57]. However, these routes can only be used after unsuccessful trans-IPS attempts [11, 56].

### Transorbital approach

Transorbital access to the CS belongs to the transvenous approach, including direct puncture of the superior and inferior OVs and CS or direct puncture of the SOV after surgical exposure [5, 15].

The superior and inferior OVJs can be punctured under sonographic and biplane roadmap guidance [58, 59]. After anterior orbitotomy, the exposed SOV can be punctured easily, with the advantage of a lower possibility of injuring the intraorbital structure

and reduced distance to the CS [5, 60]. When direct puncture or orbitotomy to access the SOV fails, a more radical method of performing orbital removal for catheterization of the SOV is another practical alternative [61].



**Figure 2. Typical case of a CS-DAVF with TVE via the IPS.** An angiogram of the right (A-B) and left (C-D) common carotid arteries in AP (A, C) and lateral (B, D) views shows a CS-DAVF supplied by the bilateral dural branches of ECA and ICA. E, The microcatheter advances to the CS via the right IPS. F, The bilateral CSs are packed by Onyx. G-H, A postprocedural angiogram of the right common carotid artery in the lateral view shows that the CS-DAVF is obliterated. **Abbreviations:** AP, anteroposterior; CS, cavernous sinus; CS-DAVF, cavernous sinus dural arteriovenous fistula; ECA, external carotid artery; ICA, internal carotid artery; IPS, inferior petrosal sinus; L: left; R: right; TVE, transvenous embolization.

Previously, transorbital CS puncture was not widely adopted due to the lack of direct or indirect visualization. Currently, the modern angiographic system has advanced greatly with the progression of software dedicated to reconstructing cone-beam computed tomography (CT). Currently, direct puncture of the confluence of the superior and inferior OVs under the guidance of planning software is safe and feasible [59].

### Other transvenous approaches

#### Puncture or incision access through other head and neck veins

In cases with the SOV as the main drainage vessel, access to the CS via the facial vein (FV) is possible [11, 62]. Matsumoto et al. (2017) reviewed previous reports of the transfemoral FV approach to the CS, and the success rate varied from 50%-100% [11]. In cases of unsuccessful transfemoral FV, direct puncture or incision of the FV can be attempted [62-64].

In addition to the FV, other veins, such as the superficial temporal vein, retromandibular vein, and jugular vein, can also be punctured to access the CS [11, 65]. To more efficiently puncture these veins, a small skin incision can be made [66]. It is helpful to create a microwire loop, which can increase the possibility of passing these abruptly angulated and tortuous veins [11, 56, 67].

#### Transcranial vein access

Rarely, transcranial puncture to the superficial middle cerebral vein can be used to perform EVT for CS-DAVF, which combines open surgery and endovascular techniques [68, 69]. Especially when the CS-DAVF has caused an intracranial hematoma, it is appropriate to evacuate the hematoma and embolize the CS-DAVF in a single session [35, 70]. In 2017, Akamatsu et al. reviewed four cases of CS-DAVFs, all of which underwent successful EVT via the superficial middle cerebral vein [70].

#### Transforamen ovale access

The approach to CS through the foramen ovale and catheterization of the skull base emissary vein can be selected as an alternative when there is no possibility of a conventional venous approach. For instance, Cabral De Andrade et al. (2012) and Gil et al. (2013) performed EVT of CS-DAVFs through the foramen ovale [71, 72].

### Transarterial embolization

When TVE is difficult or the DAVF only involves a single-side dura of the CS, it is limited in size, and with few dural feeders, TAE can be attempted [44]. If

the microcatheter tip can be wedged into the fistula with an ideal position, the TAE is safe and effective [22, 73]. The MMA and accessory meningeal artery provide gold routes to occlude the CS-DAVF [74]. In addition, in CS-DAVFs fed by APhA, TAE via APhA or combined with other routes can be a reasonable and effective choice [3, 14, 75].

During TAE, the Onyx and NBCA need to penetrate into the fistula and draining veins, as well as retrogradely into the branches of other arterial feeders [76]. In early years, NBCA was more popular in TAE for CS-DAVFs [13]. Currently, Onyx has become the first choice [44]. During TAE, to avoid embolization of dangerous anastomoses, protective balloon navigation and inflation to the cavernous ICA during injection is a feasible method [32, 77].

Two illustrative cases with TAE of CS-DAVFs are provided in Figures 3 and 4.

### Combined radiosurgery

Currently, stereotactic gamma-knife radiosurgery (SGKR) with minimal invasiveness has become an alternative primary management for DAVFs. In Hung et al.'s (2019) study, after SGKR, CS-DAVFs were more likely to be obliterated than non-CS-DAVFs [78]. Therefore, when EVT cannot obliterate the CS-DAVFs, stereotactic SGKR may be considered [5].

Using a therapeutic radiation dose of 20-50 Gy, SGKR can induce injury to the targeted vessel, thus obliterating the DAVFs. Rates of up to 90% obliteration and 85% symptomatic improvement can be achieved [79, 80].

However, the drawbacks of SGKR are continued symptoms and risk of hemorrhage in the presence of retrograde cortical drainage or eye symptoms during the latency period [80]. To overcome this limitation, a combined approach consisting of EVT and the following SGKR can be used [79].

### Prognosis and complications

In general, EVT, by way of various access routes, is a feasible and effective method to manage CS-DAVFs [27, 67, 81].

### Angiographic results and prognosis

The angiographic results of EVT can be classified into complete cure, minor residual shunt with occlusion of the superior and inferior OVs and CS, partial effect if shunt reduction was <80%, and failed attempt if there was residual filling of the superior and inferior OVs and deep venous system or cortical vein [27].

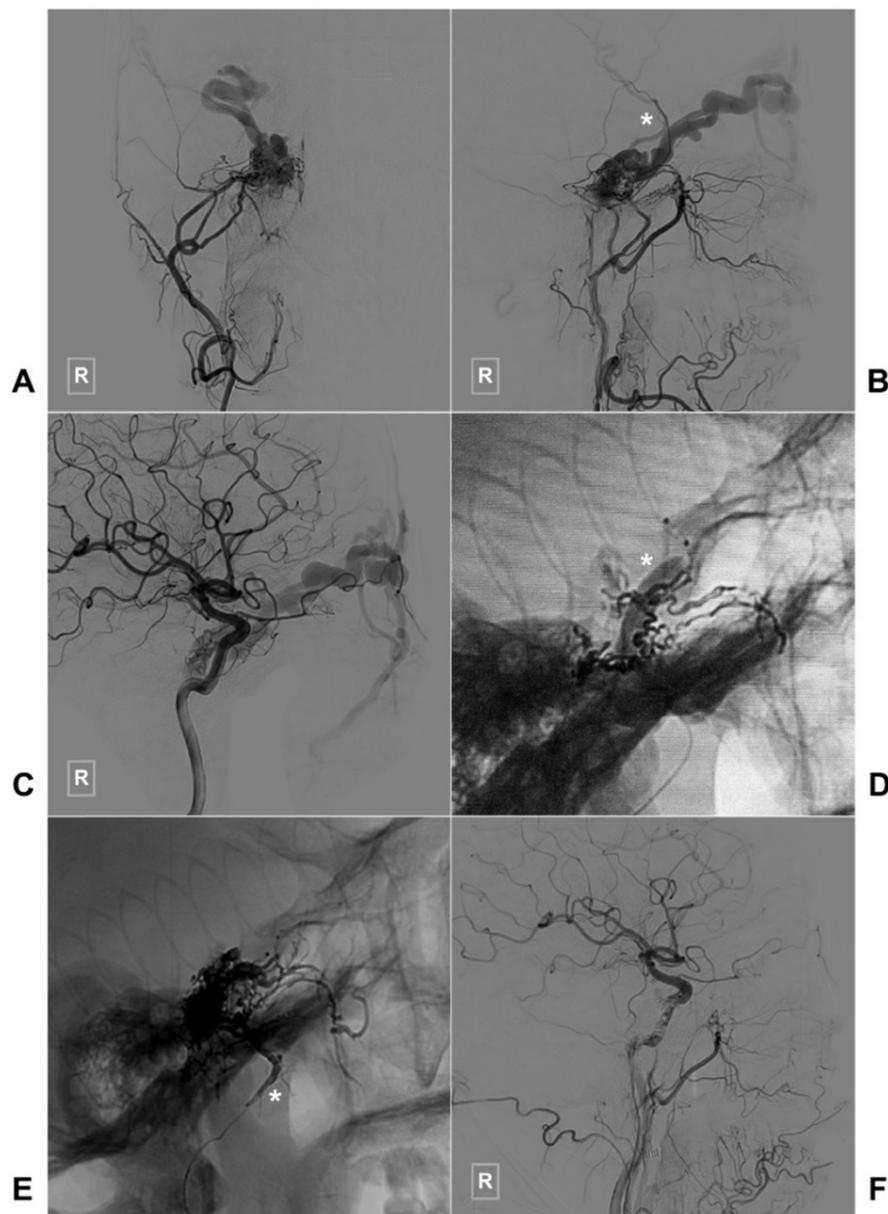
In EVT, the rate of total occlusion of CS-DAVFs varies from 51.7% to 90% using coils, from 80% to

100% using coils and NBCA, and from 76.9% to 100% using coils and Onyx [5, 13, 16, 41, 44, 48, 82, 83]. A higher rate of complete occlusion is observed using liquid agents combined with coils than using coils alone.

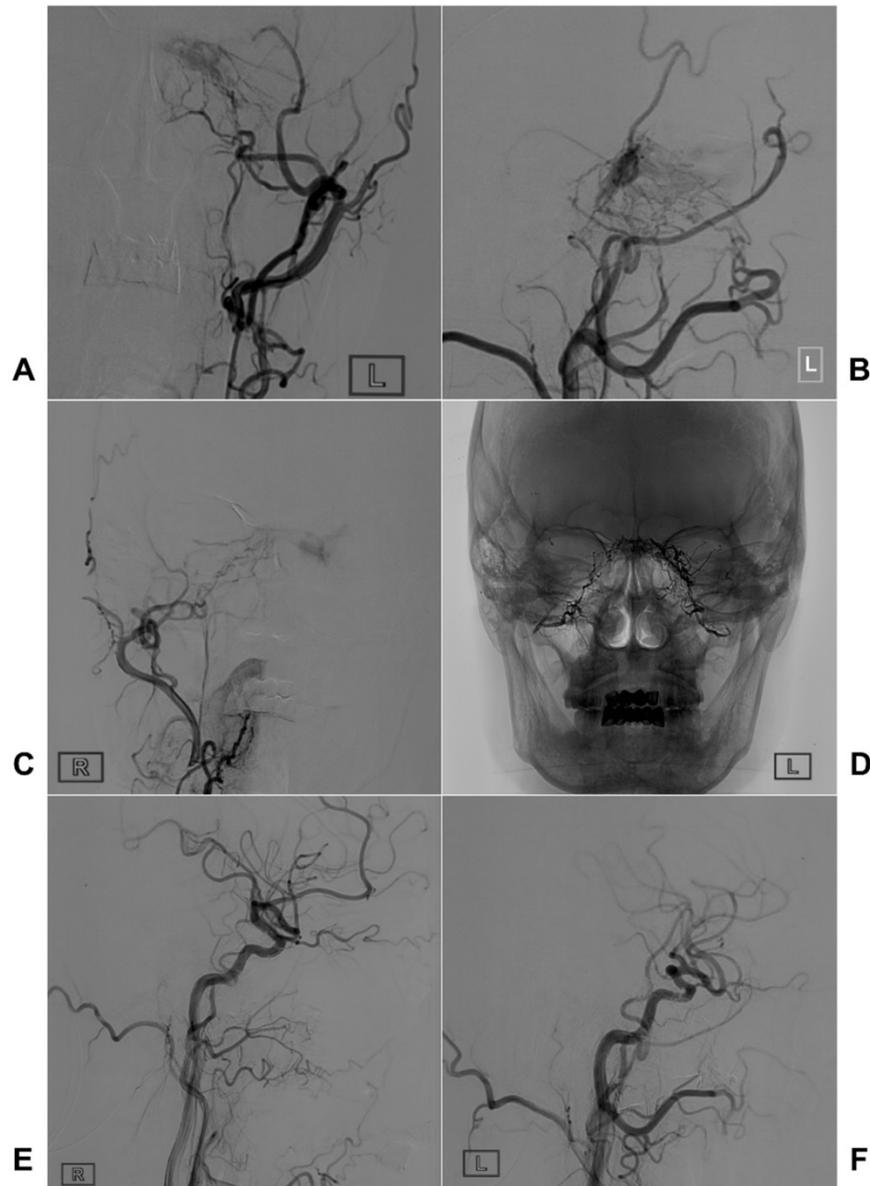
However, complete occlusion is not the ultimate goal of EVT for CS-DAVFs [83]. The primary goal should be eliminating cortical venous reflux and major retrograde shunt flow because long-term follow-up shows that the residual shunt flow can eventually disappear [83, 84]. EVT of CS-DAVFs can result in a good prognosis.

## Complications

The overall rate of complications ranges from 2.3% to 53.3% [16, 41, 48, 83]. In recent years, the rate has decreased significantly [16, 83]. In de Castro-Afonso et al.'s (2018 and 2019) studies with more than 60 patients, the complication rate was 3.2% [83, 85]. In Alexander et al.'s (2019) study with 267 patients, the complication rate was 3.6%, and permanent symptomatic complications only occurred in 0.51% of the patients [16].



**Figure 3. Typical case of a CS-DAVF with TAE via the right MMA.** An angiogram of the right ECA in AP (A) and lateral (B) views shows that the CS-DAVF is supplied by the MMA and accessory meningeal artery and drains to the superior ophthalmic vein and superficial middle cerebral vein (asterisk in B). C, An angiogram of the right ICA in the AP view shows that the dural branches of the ICA also supply blood to the CS-DAVF. D, During TAE, a balloon (asterisk) in the ICA is inflated to protect the ICA. E, An unsubtracted angiogram shows the Onyx casting via MMA (asterisk), F: An angiogram of the right common carotid artery shows that the CS-DAVF is obliterated. **Abbreviations:** ECA: external carotid artery, CS: cavernous sinus, DAVF: dural arteriovenous fistula, DSA: digital subtraction angiography, ICA: internal carotid artery, MMA, middle meningeal artery TAE: transarterial embolization.



**Figure 4. Typical case of a CS-DAVF with TAE via bilateral MMAs.** An angiogram of the left ECA in AP (A) and lateral (B) views shows that the CS-DAVF is supplied by the MMA and internal maxillary artery. C, An angiogram of the right ECA in the AP view shows that the right MMA also supplies blood to the CS-DAVF. D, An unsubtracted angiogram shows the Onyx casting via bilateral MMAs. E-F: A follow-up angiogram of the right (E) and left (F) common carotid arteries six months later shows that the CS-DAVFs are completely obliterated. **Abbreviations:** AP, anteroposterior; CS, cavernous sinus; CS-DAVF, cavernous sinus dural arteriovenous fistula; ECA, external carotid artery; MMA, middle meningeal artery; TAE, transarterial embolization.

Compared with TVE, TAE has a higher complication rate [16, 86]. In TVE, coil embolization has fewer complications than liquid embolic agents [16, 87]. The complications involved in EVT of CS-DAVFs can be divided into technical complications, cranial nerve palsies (CNPs), hemorrhagic and ischemic complications, and other complications [4].

#### Technical complications

In EVT, the microwire and microcatheter can lead to vessel injury [32]. During TVE, navigation through an occlusive IPS may cause venous injury by the sharp microwire tip, and other transvenous routes have a higher rate of complications with possible

venous rupture [67]. In transorbital approaches, the complications include supraorbital nerve and cranial nerve injuries, injury to the eyeball, infection, bleeding from the superior and inferior OVVs, and orbital or subarachnoid hemorrhage [56, 67]. Therefore, careful preoperative planning, meticulous manipulation during puncture, and vigilant postoperative management are necessary to avoid these technical complications [88].

#### Cranial nerve palsy

In TAE, CNPs can result from compromise of the blood supply of relevant cranial nerves or inadvertent nontargeted embolization of dangerous ECA-ICA

anastomoses [44, 81, 89]. In TVE, CNPs may result from the mass effect of the overpacked CS or direct injury by coils or microwire/microcatheter [15, 30, 54, 55, 90].

In all CNPs, abducens nerve palsy is the most common after TVE and can occur immediately or delayed; usually, most CNPs recover within 6 months [41, 91]. To avoid CNPs in TVE, the embolic material volume should be reduced, so superselective targeted occlusion of the SP is necessary [24, 45].

### Other complications

In TAE, the most dangerous complication with liquid embolic agents is reflux through the meningohypophyseal or inferolateral trunks into the ICA and subsequent cerebral infarction, which must be avoided [16]. Balloon protection in the ICA can be helpful.

In EVT, including TAE and TVE, due to direct toxicity from the dimethyl sulfoxide/Onyx combination, Onyx embolization of CS-DAVFs can carry the potential risk of trigeminocardiac reflex-induced bradycardia [27, 33, 92-95]. Therefore, interventional neuroradiologists should bear in mind this rare consequence prior to injection of the embolic material [92].

Rarely, EVT for CS-DAVFs can result in venous brain stem infarction, which is suspected to be caused by insufficient packing of the posterior part of the CS, which diverts the embolic agent into the deep venous and brainstem venous system. Hence, during EVT for CS-DAVFs, occluding the entrance of bridging and cortical veins is very important [48, 84, 96].

### Summary

In summary, for higher risk CS-DAVFs, EVT is the first-line option. The transvenous approach via the IPS is considered the most effective method. The transorbital approach to obtain transvenous access to the CS is another choice. For CS-DAVFs, it is usually difficult to achieve a high cure rate with TAE. With regard to the embolic materials, coil, Onyx, and NBCA can be used. Coil is the preferred material. If the EVT cannot obliterate the CS-DAVFs, SGKR may be considered after obliteration of the risk factors. Despite various complications, EVT is an effective method to treat CS-DAVFs.

### Competing Interests

The authors have declared that no competing interest exists.

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