



Spontaneous Cerebrospinal Fluid Leak at the Clivus: Two Case Reports and Literature Review

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Abstract

Introduction: Spontaneous or non-traumatic cerebrospinal fluid leaks comprise 5–10% of all csf rhinorrhea. Generally csf rhinorrhea occur at Cribriform plate, sella, sphenoid sinus and ethmoid air . Primary csf rhinorrhea from clival defect is extremely rare. We herein describe two cases of spontaneous CSF rhinorrhea through the clivus repaired with endoscopic endonasal trans sphenoid approach. Moreover, we collected evidence in the literature regarding potential etiology, symptom and treatment (which occurred in the case reported).

Background: Spontaneous cerebrospinal fluid leaks comprise 5–10% of all CSF rhinorrhea. Generally, CSF rhinorrhea occur at cribriform plate, sella, sphenoid sinus and ethmoid air. Primary CSF rhinorrhea from clival defect is extremely rare. we describe two cases of spontaneous CSF rhinorrhea through the clivus defect and review the literature.

Case Description: The first patient was a 36 -year-old female admitted to our department because of clear watery discharge from the right nostril of 3 weeks which aggravated in prone position. The second case was a 57-year-old man referred to our department with the complaint of intermittent rhinorrhea starting 6 months before surgery. He had a past history of bacterial meningitis few month before stating the rhinorrhea which was treated in another center. In both cases, testing of the fluid for beta-2 transferrin was positive. Magnetic resonance imaging and computed tomography cisternogram showed CSF leak through clivus into the sphenoid sinus. In both patients defect was repaired with abdominal fat, reinforced by fascia lata and naso septal flap via “two nostrils – four hands” endoscopic trans nasal technique.

Conclusion: At times, the exact pathophysiology of CSF clival fistula is debated, however a combination of anatomical and functional factors play a role in the occurrence of this rare phenomenon. To date, only 16 cases are reported, and the current study reported a group of two consecutive cases. To date, endoscopic trans nasal approach is the best therapeutic option to repair midline skull base defect such as the current cases.

Keywords: Spontaneous Cerebrospinal Fluid Leak; Rhinorrhea; Clivus; Meningitis; Endoscopic Endonasal Approach

Introduction

Spontaneous or non-traumatic cerebrospinal fluid leaks comprise 5%–10% of all cerebrospinal fluid (CSF) rhinorrhea [1,2]. Generally, CSF rhinorrhea can occur at the cribriform plate, sella, sphenoid sinus, or ethmoid air cells [3,4]. However, primary CSF rhinorrhea due to clival defect is extremely rare. The current study describes two cases of spontaneous CSF rhinorrhea through

the clivus which were repaired with endoscopic endonasal trans-sphenoidal approach.

Such cases are extremely rare and upon literature review, only 16 cases with clival defect are reported thus far.

The peculiar aspect of this case report is related to its rarity. Moreover, evidence was collected from the literature regarding

potential etiology, symptom, and treatment (occurred in the reported cases).

Case 1

A 36 -year-old female was admitted for three weeks of clear watery discharge from the right nostril, which was aggravated in prone position. The patient denied any recent trauma. A review of systems was negative except for headaches and nasal discharge. The nasal fluid tested positive for beta-2 transferrin, indicating that the fluid was CSF. Brain MRI revealed that the sphenoid sinus

was filled with cerebrospinal fluid (CSF) and sagittal T2 weighted MRI revealed a fistula tract from prepontine cistern to sphenoid sinus (Figure 1). There was no evidence of benign intracranial hypertension. Computed tomography cisternography revealed that the contrast material passed from the prepontine cistern into the sphenoid sinus through this bone defect in the clivus (Figure 1). Before surgery, a lumbar puncture was performed to administer 0.25 mL of 10% fluorescein with 10mL of cerebrospinal fluid (CSF) to help visualize CSF leaks during surgery and to ensure there was no leak after reconstruction of the defect.

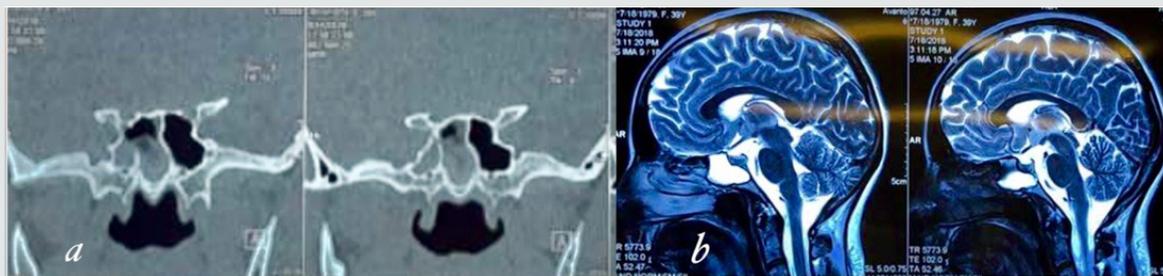


Figure 1: (a) Coronal CT cisternography shows sphenoid sinus filled with CSF.(b) Sagittal T2-weighted MRI of the brain showing cerebrospinal fluid (CSF) leak into the sphenoid sinus through clival defect.

The patient underwent endoscopic trans nasal trans sphenoid surgery. The anterior and middle portions of the clivus were exposed between both carotid arteries. During surgery, the defect was defined to the left of the midline in the clivus. The basilar artery

was seen through the defect in prepontine cistern (Figure 2). The defect was closed with a multilayer reconstruction consisting of fat, fascia lata, and naso septal flap (Figure 3). There was no recurrence of CSF leak at 2 years follow-up.

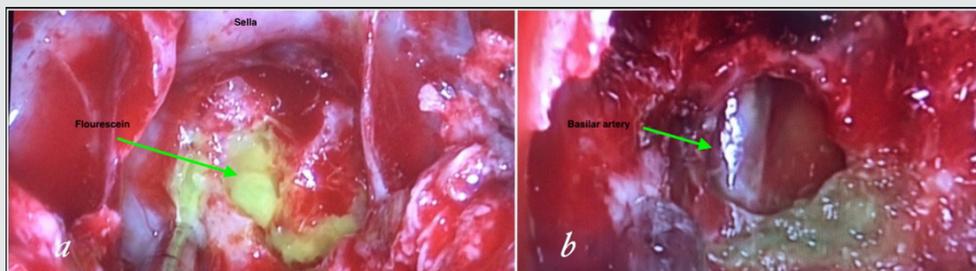


Figure 2: (a) Intraoperative endoscopic view show CSF leakage from clivus defect just inferior to the sella . (b) Basilar artery in prepontine cistern behind the defect.

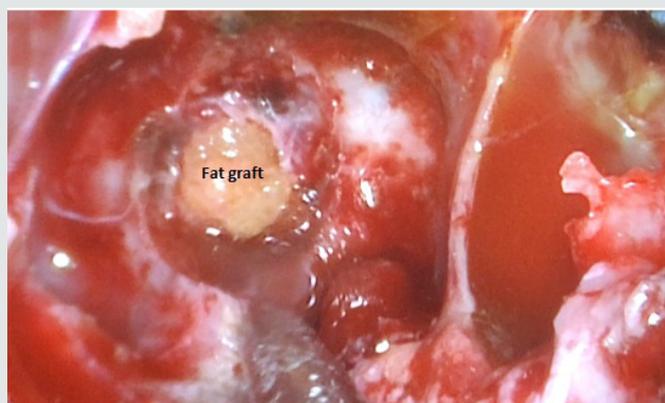


Figure 3: Endoscopic view after repair of the defect with abdominal fat.

Case 2

A 57-year-old man referred for clear watery discharge from the right nostril of no obvious cause. He suffered from intermittent rhinorrhea starting 6 months prior to arrival. He reported recent history of bacterial meningitis one month ago, which was treated successfully at an outside hospital. On admission, he had no focal neurological deficits. Nasal fluid tested positive for beta2 transferrin. Brain MRI revealed that the right sphenoid sinus was filled with CSF (Figure 4). CT cisternography showed that the contrast material passed from the prepontine cistern into the

sphenoid sinus through this bone defect in the clivus (Figure 4). After intrathecal administration of 0.25 mL of 10% fluorescein with 10 mL of cerebrospinal fluid the patient underwent endoscopic trans nasal approach. After stripping the mucosa from posterior wall of sphenoid sinus, CSF leak was observed in the upper region of clivus just below the sella at the midline (Figure 5). The defect was closed by abdominal fat and reinforced by fascia lata and naso septal flap. At the During 30-month follow-up appointment, no signs of recurrence were found. In both of the above reviewed cases, a lumbar drain was not placed pre or postoperatively.

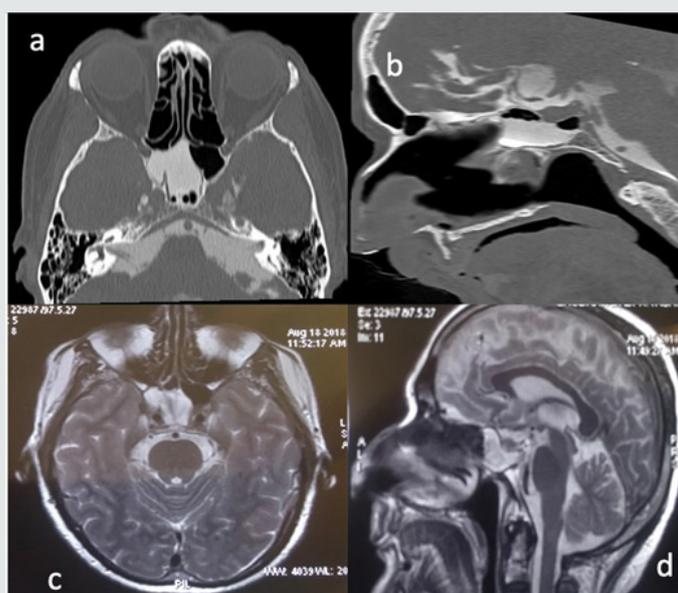


Figure 4: (a) CT cisternography showing right sphenoid sinus filled with CSF and (b) the entry of cerebrospinal fluid into the sphenoid sinus. (c) Axial T 2 weight MRI revealed right sphenoid sinus filled with CSF. (d) Sagittal T2 MRI shows CSF leakage from prepontine cistern to sphenoid sinus.

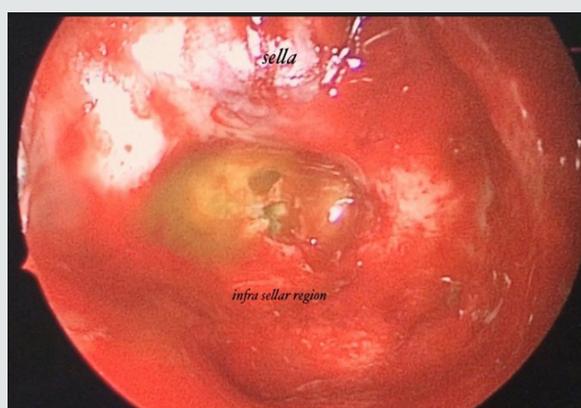


Figure 5: Intraoperative endoscopic view shows CSF leakage from clival defect.

Discussion

Cerebrospinal fluid (CSF) leaks most commonly result from nonsurgical trauma (80%-90% of cases), followed by surgical procedures (16%), and nontraumatic or spontaneous causes (4%)

[5,6]. O'Connell first subcategorized spontaneous CSF rhinorrhea in two groups in 1964; primary spontaneous, when there is no cause for skull defect, and secondary spontaneous, when a cause can be found [7,8] Defects in the roof of the ethmoid sinus or in the floor

of the anterior cranial fossa contribute to the most common site of fistula in the patient with traumatic CSF rhinorrhea. However, for primary spontaneous CSF fistulas, a sphenoidal fistula is most common (60%).

In these cases the junction of the floor of the middle cranial fossa to the lateral wall of the sphenoid sinus is the most common site of CSF leak through sphenoid sinus, [3] however, primary spontaneous CSF leaks from clival defect are extremely rare. In a study by Hooper on 138 sphenoid bones, the defect of bones connecting the sphenoid to the cranium leading to CSF leak was observed in 5% of cases; all of which were in lateral wall of the sphenoid sinus [9,10]. To the best of the authors knowledge, only 16 cases of spontaneous CSF leaks from clival defect are reported thus far. The exact pathophysiology of CSF clival fistula is debated. Morley and Wortzman postulated that congenital bone defects in the middle fossa can explain the leaks through the lateral extensions of the sphenoid sinus [11]. However, from anatomic point of view, there is no clear embryological evidence to explain clival defect causing CSF rhinorrhea.

The clivus is a bony structure composed of the fusion of the posterior portion of the sphenoid body (basisphenoid) and the basilar part of the occipital bone (basioccipital) at the sphenoccipital synchondrosis (SOS). SOS fusion can occur at any age, but it usually happens before adolescence. SOS may persist into adult life and may be mistaken for a fracture or defect.

However, this synchondrosis is caudal to the future sphenoid sinus; otherwise, clival ossification would be a continuous enchorial ossification without fusion point that could explain bone defect [12,13]. According to Faizuddin Ahmad et al. most authors believe that excessive pneumatization of sphenoid sinus causes a thin bony wall at some points of clivus and sphenoid. These phenomena, combined with other potential factors such as arterial pulsation and continuous CSF pressure wave ultimately lead to the bone defect at the clivus and CSF leak [7,14,15]. Sixteen cases of spontaneous CSF leak at the clivus were reported from 1995 to 2018 (Table1). In all patients, the defect was localized in the upper clivus. In all sixteen cases reported thus far CSF leak from clival defect occurred in adult patients, which may explain the role of CSF pressure pulsation as a predisposing factor. The CSF pressure pulsation reaches its maximum point in adults, approximately three times higher than that of infants. Eleven of the sixteen patients were female. In 1995, Coitero et al., reported the first two cases of CSF fistula through the clivus. In one case they suggested that basilar artery pulsations over a thinned bone structure could be the cause of clival defect [16]. Radiographic features of increased intracranial pressure (ICP) such as empty sella syndrome (80%) and arachnoid pits (63%) are often observed in patients with spontaneous CSF rhinorrhea. However, in a case report by Thibaut van Zele in spontaneous clival CSF leak, radiologic signs of increased ICP (empty sella and/or arachnoid pits) were observed only in two cases of four patients [17].

Table 1: Published Cases of clival defect leading spontaneous csf leak.

Case Reference	Sex	Age	Meningitis	Duration of cdf Rinorrhea	Previous surgery with no finding defect	Surgical approach for defect closure	Recurrent
Guang Yong Chen 2018(18)	MALE	71	-	3 week	NO	ETSS	NO
Vivek Tandon 2017(24)	female	55	+	3 week	YES (ETSS)	sublabial rhino septaltranssphenoidal	NO
Sheikh Asad 2017 (10)	female	64	+	2 y	NO	ETSS	YES
Satiawaty Mohindra 2017 (28)	female	35	-	2 m	YES (ETSS)	ETSS	NO
Krzysztof Oleś 2015(25)	female	60	+	3 y	NO	ETSS	NO
Thibaut Van Zele 2013 (17)	Female	37	-	2 m	NO	ETSS	NO
	Female	61	-	4m	NO	ETSS	NO
	Female	78	-	3m	NO	ETSS	NO
	Female	42	-	2m	NO	ETSS	NO
	Female	48	-	3m	NO	ETSS	NO
	Male	50	-	1m	NO	ETSS	NO
Luana Antunes Maranhã 2012 (27)	female	49	+	10 y	YES (3time)	ETSS	NO
Faiz Uddin Ahmad 2006 (7)	Male	52	+	-	NO	ETSS	NO
	male	56	-	2 y	NO	sublabial rhinoseptal trans-sphenoidal	NO
Coiteiro1995(16)	Female	53	+	3 m	NO	ETSS	NO
	male	43	+	1 week	NO	ETSS	NO

In all of the current study patients, the CSF pressure was normal and magnetic resonance findings were not suggestive of benign intracranial hypertension. Radiologic evaluation of CSF leaks is a diagnostic challenge that often involves multiple imaging studies. Various imaging studies such as contrast-enhanced computed tomography (CT) cisternography, radionuclide cisternography, and MR cisternography are employed in the diagnosis of such cases. To date, contrast-enhanced CT cisternography is the standard for CSF leak evaluation with a sensitivity of 76-100% [18,19].

However, cisternography requires intrathecal injection via lumbar puncture which can cause discomfort for patients. Additionally, CSF leak is often an intermittent phenomenon, therefore the sensitivity of cisternography depends on the timing of the examination. Additional MRI (with T2-weighted sequences) can be helpful if parenchymal or meningeal herniation is suspected [20,21]. Of the above referenced two case studies, one patient's sphenoid sinus was filled with cerebrospinal fluid and defect was discovered in the upper part of the clivus.

Retrospectively, in cases suspected of clival defect, the T2-weighted sequence of mid-sagittal MRI showed CSF fistula from prepontine cisterna to sphenoid sinus. Several surgical approaches can be utilized to repair CSF rhinorrhea. However, to date endoscopic trans nasal trans sphenoid surgery is established as the standard technique to repair CSF leak. This approach is less invasive, with lower morbidity and mortality, excellent view of the surgical field, and a higher success rate [22-25]. In the current study, endoscopic trans nasal approach was performed for fistula repair in both patients. All defects were closed with a combination of fat, fascia, and a free or nasoseptal flap. Review of the literature highlighted the fact that identification of the bone/dural defect responsible for CSF leakage was the most important point for successful surgical intervention, since a missed site can lead to improper treatment and recurrence of the leak [25,26].

In a report by Vivek Tandon, sphenoid sinus packing was performed when the exact site of the leak was not identified, and the patient presented with recurrent rhinorrhea and meningitis [24]. In a report by Luana Antunes Maranhã, the patient underwent anterior skull base repair via bifrontal craniotomy three times since the exact site of leakage was not defined and the patient presented with recurrent CSF leak. Finally, the patient underwent endoscopic trans nasal approach for clival defect closure [27]. In some cases, there is more than one defect that should be repaired. For example, in a study by Satiawaty Mohindra, a patient had two concomitant defects. In the first surgery, clival defect was missed; the right cribriform plate and right sphenoid sinus defect closures were performed. CSF rhinorrhea recurred after one month and the patient underwent revision surgery. In revision surgery, previous repair site did not show leakage, but a high-pressure leak was observed through posterior wall of sphenoid sinus from clivus [28].

In all of the current study patients, clival defect was the only defect causing CSF leak. To date, various materials such as mucoperichondrium, cartilage, fat, fascia, and fibrin glue have been utilized to seal the fistula with different success rates. Fat and fascia were used for defect closure in all of the patients in the current study, comparable to patients with spontaneous CSF leak or repair of dural defect after adenoma surgery. Similar to Hegazy, we believe that the material used in the closure of the fistula is not important in the success of the intervention. The important key to success is the determination of the bony edge surrounding the defect [29].

Similar to the other authors, we reserved lumbar drainage for patients with elevated intracranial pressure. In both patients, the conservative measures such as bed rest, elevation of the head, and avoidance of straining activities were implemented after surgery [30,31].

Conclusion

Spontaneous CSF rhinorrhea located at the clivus is an extremely rare condition. To date, only 16 cases are reported, and the current study reported a group of two consecutive cases. It seems that a combination of anatomical and functional factors play a role in the occurrence of this rare phenomenon. To date, endoscopic trans nasal approach is the best therapeutic option to repair midline skull base defect such as the current cases.

Here, the initial outcome was successful, but long-term follow-up is required.

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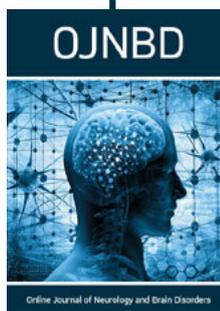
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