ABSTRACT
Cerebrospinal fluid rhinorrhea results from a breakdown of barriers between the subarachnoid space and aerodigestive tract. Because of the risk of potential intracranial complications, all persistent cerebrospinal fluid (CSF) leaks should be repaired. Over the past few decades, with the advances in endoscopic techniques, the surgical management of CSF rhinorrhea has changed significantly. CSF leak repair has advanced from the conventional open intracranial approaches by neurosurgeons to the transnasal endoscopic ones by otorhinolaryngologists. In this study, 35 patients with CSF leaks underwent endoscopic repair with a 94.3% first attempt success rate, and 100% overall success rate. There were no major intraoperative or postoperative complications. To conclude, the endoscopic approach has a high success rate with minimal morbidity and has therefore, become the preferred method.

Keywords: Skull base, CSF rhinorrhea, Repair, Endoscopic.

INTRODUCTION
Cerebrospinal fluid (CSF) rhinorrhea results from an abnormal communication between subarachnoid space and nose due to breakdown of all the barriers between them, namely, arachnoid membrane, dura mater, bone of skull base and the mucosa of nasal cavity and paranasal sinuses. It can be traumatic or nontraumatic in origin. The incidence of CSF rhinorrhea has increased greatly as a result of the alarming rise of road traffic accidents and advanced endoscopic skull base surgeries. The commonest cause of CSF leaks is accidental trauma, followed by iatrogenic trauma and non-traumatic spontaneous leaks. Most of the cases respond to conservative management, that includes— bed rest with reverse — Trendelenberg position, medical treatment and placement of lumbar drain. Any persistent CSF leak must be surgically repaired to avoid imminent life-threatening complications, like meningitis, brain abscess and pneumocephalus.

Galen, in the 2nd century AD, first described the leakage of CSF after cranial trauma, but it was not considered a pathologic process until the 17th century. In 1826, Charles Miller gave the first comprehensive description of CSF leak in a child with hydrocephalus. In 1899, Sir St Clair Thomson, reported a series of spontaneous nasal CSF leaks and coined the term ‘CSF rhinorrhea’. The history of surgical repair of CSF rhinorrhea began in 1926 when, Walter Dandy, an American surgeon reported the first successful intracranial repair using a bifrontal craniotomy. This procedure has a success rate of 60 to 80%. After nearly two decades, Gosta Dohlman, a Swedish physician, performed the first extracranial approach for the repair of a CSF leak from the roof of the ethmoid using a naso-orbital incision. In 1952, Oskar Hirsch was the first to repair a CSF leak in the sphenoid sinus using the trans-septal approach. Wigand, in 1981, performed the first successful transnasal endoscopic approach to repair anterior cranial base CSF rhinorrhea. Once considered the province of neurosurgeons, CSF leak repair was primarily managed by open intracranial approach. Over the past few decades, as a result of advances in endoscopic tools and techniques, the surgical management has changed significantly, and now, CSF leaks are being approached extracranially by minimally invasive endoscopic techniques by otolaryngologists. These approaches are associated with minimal morbidity and mortality, and over 90% success rate.

The aim of this study was to evaluate the outcomes in patients who underwent endoscopic repair of CSF rhinorrhea and to share our experience.

PATIENTS AND METHODS
This was a prospective study carried out in the Department of Otolaryngology and Head and Neck Surgery, Sawai Man Singh Medical College and Hospital, Jaipur, a tertiary level referral hospital. From November 2009 to December 2012, 35 patients who presented with CSF rhinorrhea were included.
Detailed history was taken including history of rhinorrhea, history of trauma/previous surgery, recurrent meningitis. Diagnosis was ascertained by nasal fluid analysis for glucose, electrolytes, cytology and proteins. Diagnostic rigid nasal endoscopy was done in all the patients. All patients underwent CT cisternography using metrizamide dye to localize the site of defect. MRI cisternography (Fig. 1) was reserved for the cases where the site of leak could not be localized by CT.

Transnasal endoscopic repair of CSF leak was done in all the patients under hypotensive anesthesia, using 4 mm 0° and 30° rigid nasal endoscopes. The location of fistula was identified using the nasal endoscope and the margins of the defect were defined. Nasal or sinus mucosa surrounding the defect was removed for around 5 mm to expose the bone around the defect, thereby allowing attachment of the graft to the bone. Cephaloceles, if present, were managed by removing the mucosa from the mass and reducing the mass through the bony defect, by direct manipulation or by bipolar cautery (Fig. 2). Depending on the site and size of defect, fistula closure was done by underlay-overlay technique or bath plug technique. Underlay-overlay technique was preferred for larger defects, in which repair was done in five layers. Adipose tissue and fascia lata were harvested from the thigh using a separate incision. The first layer was fat plugging the defect. Fascia lata was then tucked in by underlay technique between the dura and skull base. This was followed by a layer of microfibrillar collagen. The fourth layer was fascia lata placed by overlay technique, over the dural lesion and over the exposed bony margins, which were denuded of mucosa. Another layer of microfibrillar collagen was placed over this. Graft was stabilized using oxidized cellulose and absorbable gelatine sponge. Bathplug technique (Figs 3A and B) was used to seal small defects. In this method, adipose tissue was harvested from thigh/ear lobule. Fat was placed over the defect and incorporated intracranially. This was held in place by the ambient intracranial pressure as well as a soft tissue graft (fascia lata) centered over the defect with the edges of the centered graft exceeding the bony defect circumferentially. Next, a layer of microfibrillar collagen was placed over it. Graft was stabilized using oxidized cellulose and absorbable gelatine sponge.

Fig. 1: MR cisternogram 5 mm defect seen in left cribriform plate with CSF leak through it and herniation of gyrus rectus also seen

Fig. 2: Endoscopic image of a meningoencephalocele (arrow) from the cribriform plate (left side)

Figs 3A and B: Endoscopic image of fat being obliterated into the site of defect by bathplug technique
Following surgery, all patients were advised strict bed rest with head elevation (30°), minimal exertion and no straining. Postoperatively, all patients were given i.v. antibiotics — 3rd generation cephalosporin, aminoglycosides and anaerobic coverage, i.v. mannitol, oral acetazolamide, cough suppressant and stool softeners. Nasal pack was removed 5th to 7th postoperative day. The patients were advised not to strain and lift heavy weights for at least 1 month and were prescribed cough suppressants and stool softeners. Follow-up was done by nasal endoscopy every fortnight for 2 months and then monthly to see for any recurrence (Fig. 4). The follow-up period varied from 6 to 24 months.

RESULTS

Among the 35 patients who were included in the study, there were 20 males and 15 females. Age group varied from 10 to 60 years with mean age of 35 years. CT cisternography indicated the site of leak in all but three cases in which MR cisternography was required. Five patients had coexistent cephaloceles. Two patients had history of recurrent meningitis which was managed conservatively and following its resolution, these patients underwent endoscopic repair of CSF leak. None of the patients had bilateral leaks or prior attempted repair.

The most frequent site of cranionasal fistula was cribriform plate (20/35; 57%), followed by roof of ethmoids (10/35; 29%), sphenoid sinus (4/35; 11%) and frontal sinus (1/35; 3%).

From the etiological point of view, dural defects were classified into three groups:

- Spontaneous/nontraumatic leaks (20/35; 57%) — this group included patients with no history of trauma or previous surgery.
- Surgical/iatrogenic trauma (7/35; 20%) — these patients had history of previous nasal or skull base surgery: pituitary surgery in four patients, surgery for ethesioneuroblastoma in two patients and sinonasal procedure in one patient.
- Accidental trauma (8/35; 23%) — these patients had history of head injury due to road traffic accident or fall from height.

The closure technique depended on the location and size of the defect. The size of the defect varied from 2 mm to 1 cm. We used the underlay-overlay technique in 28 (80%) patients and the bathplug technique in 7 (20%) patients (Table 1). None of the patients had any significant intra or postoperative complications. Three patients complained of postoperative headache which resolved within a few days.

CSF leak repair was successful at first attempt in 33/35 (94.3%) patients, and an overall success rate, including revision of 100%. Two patients presented with recurrence, out of which one had minimal leak which improved with only conservative management. The second patient underwent revision surgery and on follow-up the defect had completely resolved. The follow-up period varied from 6 to 24 months. The defect size and choice of surgical technique did not appear to influence the repair outcome.

DISCUSSION

Transnasal endoscopic repair of cerebrospinal fluid leaks is a major challenge for otorhinolaryngologists and skull base surgeons. The advances in technology have permitted its repair by the minimally invasive endoscopic technique. This approach is cost-effective, less time consuming, has minimal morbidity and mortality, and a very high success rate as compared to the open intracranial approach.

The etiology of CSF rhinorrhea can be traumatic or nontraumatic. The commonest cause of CSF rhinorrhea is trauma-accidental followed by iatrogenic. The anterior skull base is extremely prone to trauma both accidental and iatrogenic.

<table>
<thead>
<tr>
<th>Table 1: Characteristics of cerebrospinal fluid leaks</th>
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<tr>
<td><strong>Etiology</strong></td>
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<td>Spontaneous</td>
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<tr>
<td>Iatrogenic trauma</td>
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<td>Accidental trauma</td>
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<td><strong>Site</strong></td>
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<tr>
<td>Cribriform plate</td>
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<tr>
<td>Roof of ethmoids</td>
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<td>Sphenoid</td>
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<td>Frontal</td>
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<td><strong>Surgical technique</strong></td>
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<td>Underlay-overlay</td>
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<td>Bathplug</td>
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</table>
In our study, there were ten patients with defect in roof of ethmoids, four in sphenoid sinus and one with frontal sinus defect. We did not use intrathecal fluorescein to identify site of leak because of suspected neurological complications.

All patients underwent endoscopic repair of CSF leak with a 94.3% success rate at first attempt and the overall success rate of CSF leak repair, including revision, was 100%. This was in concordance with the results obtained in other studies8–15 (Table 2). The defect was repaired, depending on the size, by underlay-overlay technique or bathplug technique. The high success rate was attributed to the multi-layered repair using fascia lata, adipose tissue, microfibrillar collagen and oxidized cellulose. Various other graft materials have been used in other studies to repair the defect—nasal cartilage and mucoperichondrium,8–10 middle turbinate flap,9,11,13,14 fibrin glue.8,13,14 We did not use any of these in our study. Several authors recommend the use of postoperative lumbar drain. However, in our study, we did not find it necessary and lumbar drain was not used in any of the patients. The patients were followed up for 6 to 24 months. It was found that the size of the defect and choice of surgical technique did not affect the outcome.

CONCLUSION

Transnasal endoscopic repair of CSF rhinorrhea is a safe and effective method, with minimal morbidity and as such

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**Table 2: Review of literature**

<table>
<thead>
<tr>
<th>References</th>
<th>No. of patients</th>
<th>Etiology</th>
<th>Site of defect</th>
<th>Graft material/surgical technique</th>
<th>Results and follow-up</th>
</tr>
</thead>
<tbody>
<tr>
<td>James A Burns et al13 (1996)</td>
<td>42</td>
<td>Spontaneous—10 Iatrogenic trauma—27</td>
<td>Sphenoid sinus—20 Fovea ethmoidalis—15 Cribriform plate—7</td>
<td>Sphenoid fistulae-adipose tissue with attached dermis or fascia with fibrin glue Fovea ethmoidalis and cribriform plate fistulae-middle turbinate graft with fibrin glue</td>
<td>Success rate—90.4% (38/42)</td>
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<tr>
<td>M Cassano et al12 (2009)</td>
<td>125</td>
<td>Spontaneous—43 Iatrogenic trauma—29</td>
<td>Sphenoid sinus—43 Anterior ethmoid—21 Anterior ethmoid—17 Frontal sinus—2</td>
<td>Overlay apposition of a lower turbinated mucoperiosteal flap fixated with fibrin glue and surgical</td>
<td>First attempt success rate—94.4% (118/125)</td>
</tr>
<tr>
<td>Present study</td>
<td>35</td>
<td>Spontaneous—20 Iatrogenic trauma—7</td>
<td>Cribriform plate—20 Roof of ethmoids—10 Sphenoid sinus—4 Frontal sinus—1</td>
<td>Underlay-overlay — 28/35 Bathplug technique — 7/35</td>
<td>First attempt success rate—94.3% (33/35)</td>
</tr>
</tbody>
</table>

The dura in this region is tightly adherent to underlying skull, and trauma can result in CSF rhinorrhea. Most of these leaks subside with conservative management and surgery is not required. Dural defects though rare, must be closed regardless of the cause due to potential risk of life threatening complications like meningitis and pneumocephalus.

In our study, a diverse age group of patients from 10 to 60 years underwent endoscopic repair of CSF leaks. Males were predominantly affected, with a male to female ratio of 1.3:1, in concordance with other studies.8,11,12 Diagnosis was by history of intermittent rhinorrhea, nasal fluid cytological and biochemical analysis and rigid nasal endoscopy. Fluid analysis for β2 transferrin or B trace protein is highly sensitive, however, these tests were not available in our clinical set up. CT scan was effective in localizing site of defect in most cases. MRI being more sensitive, was used for detection of site of leak when CT was inconclusive and in cases of meningoencephalocele. Two of the patients had history of recurrent meningitis. None of the patients had pneumocephalus, bilateral leaks or prior attempted repair.

The commonest cause of CSF rhinorrhea in our study was spontaneous leaks, followed by accidental trauma and iatrogenic causes. This was in contrast to other studies,8,9,12 where trauma was the most common cause. Cribriform plate was the predominant site of leak, as found in literature.9,10 In our study, there were ten patients with defect in roof of ethmoids, four in sphenoid sinus and one with frontal sinus defect. We did not use intrathecal fluorescein to identify site of leak because of suspected neurological complications.

The commonest cause of CSF rhinorrhea in our study was spontaneous leaks, followed by accidental trauma and iatrogenic causes. This was in contrast to other studies,8,9,12 where trauma was the most common cause. Cribriform plate was the predominant site of leak, as found in literature.9,10
no mortality. A multilayered technique using fat, fascia, microfibrillar collagen and oxidized cellulose to repair the defect has a high success rate. Therefore, transnasal endoscopic technique has become the preferred method for repair of cranionasal fistula.

REFERENCES