Abstract

Mucoceles of the paranasal sinus are benign expanding lesions that primarily occur between the fourth and seventh decades of life. Sinus mucoceles may develop due to the obstruction of the normal passageway by trauma, inflammation, masses, lesions, idiopathic, or iatrogenic causes. The most common site for the evolution of a mucocele is within the frontal sinuses, followed in frequency by the ethmoidal, maxillary, and sphenoid sinuses. The symptoms depend on the location and size of the mucocele. The symptoms can be classified as nasal, ophthalmologic, intracranial, and cosmetic. A mucocele can become infected, forming a mucopyoce- le, with a risk of infectious complications including meningitis, orbital cellulitis, and osteomyelitis. Today, the first choice of treatment is marsupialization of the mucoceles using endoscopic sinus surgery. External approaches to frontal sinus are still used, in erosion of the anterior frontal sinus wall and far lateral located lesions.

Keywords
Mucocele; FESS; External Approach
The paranasal sinus mucocele is a dilated mucous-filled sinus that is lined by mucous membranes [1]. Mucoceles of the paranasal sinuses are benign, expanding lesions that primarily occur between the fourth and seventh decades of life. The incidence is similar for both genders [2-4]. Paranasal sinus mucoceles are extremely rare in children and adolescents. Cystic fibrosis is the main etiologic factor [5]. The symptoms depend on the location and size of the mucocele.

The mechanism underlying mucocele formation is multifactorial. Impediments to sinus ostium ventilation are thought to be the principal cause. Sinus mucoceles may develop due to the obstruction of the normal passageway by trauma, inflammation, mass lesions, idiopathic, or iatrogenic causes [6-8] (Figure I). However, the predisposing etiology of primary and secondary nasal mucoceles seems to be different. For the formation of primary mucoceles, inflammatory blockages of mucous drainage, secretory duct obstruction, cystic dilation of mucosal glands, and cystic degeneration of polyps were proposed as possible causes. On the other hand, secondary mucoceles sequestering residual mucosa in the wound and long-term retention of tissue fluid have also been advocated as factors leading to their formation [9-11].

Bockmühl U et al. has evaluated 290 patients with sinus mucocele according to their etiologies. They reported incidences related to: a previous operation 66% (168 patients), trauma 14% (37 patients), chronic sinusitis 2% (5 patients), tumor 1% (2 patients), and unknown causes 17% (43 patients). When mucoceles due to a previous operation were further evaluated, it was concluded that 98% were due to external procedures such as Jansen-Ritter and Caldwell-Luc, and 2% were due to endonasal surgery. On average, the periods of time between external surgery and occurrence were as follows: maxillary sinus 15 years, fronto-ethmoid 13 years, ethmoid sinus 10 years, and frontal and sphenoid sinus 8 years [12,13]. It is reported that the use of alloplastic materials in the management of frontal sinus fractures may be a cause of orbital mucopyocele [14].

Mucoceles grow slowly. As pressure builds due to continuous swelling, the surrounding bony structures begin to thin, contributing to further resorption and remodeling of the bone. Some authors suggest that release of osteolytic factors may contribute to the progressive development of mucoceles or mucopyoceles. Prostaglandins and collagenases aid in osteolysis and further enhance the expansile nature of the cysts [15-17]. Lund and Milroy proposed that the obstruction to sinus outflow in combination with superimposed infection caused the release of cytokines from lymphocytes and monocytes. The cytokine release stimulates fibroblasts to secrete proteoglycans and collagenases, which in turn could stimulate bone resorption leading to expansion of the mucocele [18].

Histopathologically, paranasal sinus mucoceles have features of respiratory mucosa with cyst walls demonstrating single-layered, pseudostratified, ciliated, columnar epithelium [16,18,19]. Although metaplastic changes are rare, chronic cases show evidence of squamous metaplasia. Reactive bone formation is also possible in areas adjacent to cyst epithelium [16]. Mucoceles show elevated expression of IL-12 which secondarily leads to increased expression of IL-2 and interferon gamma. Subsequently, an increased activation of TH2 lymphocytes hastens the presence of a chronic inflammatory infiltrate [17]. Additional findings include cholesterol granuloma, hemorrhage, fibrosis, and granulation tissue [20]. Central nervous tissue may also be identified in those cases of intracranial herniation [16]. In some cases, ruptured salivary ducts may be identified. The adjacent minor salivary glands often contain a chronic inflammatory cell infiltrate and dilated ducts. Most mucoceles are self-limiting. They rupture and resolve on their own. However, other lesions are chronic in nature and local surgical excision is necessary. To minimize the risk of recurrence, when the area is excised, the surgeon should remove any adjacent salivary glands that may be supplying the lesion. The specimen should be submitted for pathologic examination to confirm the diagnosis and to rule out the possibility of a salivary gland malignancy. The prognosis is excellent, although occasionally mucoceles recur, necessitating reexcision, especially if the associated glands had not been removed [19].

Radiographically, the sinus is initially opacified by retained secretions, which displaces all the air content. Pressure erosion of the surrounding bony walls by the mucocele decalcifies the mucoperiosteal cortical plate and destroys the normal scalloped borders of the sinus (Figure II). With progression of the lesion, the sinus appears more radiolucent radiographically. There may be a zone of reactive osteitis and bone sclerosis around the margins of the mucocele. In approximately 5% of mucoceles, peripheral calcification is evident radiographically. This calcification may be large enough to stimulate the growth of an osteoma. With extensive intracranial extension by a frontal mucocele, a segment of the posterior wall of the frontal sinus may

Figure I. Frontal sinus mucocele due to nasal polyposis
be displaced as a bony fragment deep into the frontal lobe. A view of the optic foramen may be valuable in the diagnosis of a mucocele of the frontal sinus as well as of the posterior ethmoidal cells. The sphenoid mucocele presents radiographically as a destructive expanding cystic or mass lesion. When expansion and bone destruction are present, differential diagnosis includes benign and malignant lesions of the paranasal sinuses. Benign lesions include neurofibroma; dermoid, epidermoid, cementifying fibroma, angiofibroma, inverting papilloma, and cylindroma. Malignant lesions include adenoid cystic carcinoma, plasmocytoma, embryonal rhabdomyosarcoma, lymphoma, schwannoma, and tumors of dental origin [19,21,22]. In the absence of bone erosion, mucoceles must be differentiated from several conditions including retention cysts, chronic sinusitis, antrochoanal polyp, and polyposis of the paranasal cavities [23-25].

A mucocele can become infected, forming a mucopyocele with the risk of infectious complications including meningitis, orbital cellulitis, and osteomyelitis. When the mucocele expands so that the sinus volume is insufficient to house the mucocele, pain and pressure erosion can occur [26]. Pressure erosion can result in significant anatomic deformity including facial asymmetry and dynamic proptosis of the orbits [27]. Fortunately, after the resolution of the mucocele, the deformity usually remodels. Symptoms vary depending on the location of the mucocele and may be classified as rhinological, neurologic, or most frequently, ophthalmologic [28,29]. The most common site for the evolution of a mucocele is within the frontal sinuses [25,30]. Sphenoid sinus mucocele is a rare condition which represents only 1-2% of all paranasal sinus mucoceles [10]. In a study of 108 patients, Har-El G reported paranasal sinus mucocele occurrence rates as: frontal and fronto-ethmoid 66 patients (61%), ethmoid sinus 23 patients (22%), sphenoid sinus 12 patients (11%), and maxillary sinus 6 patients (6%) [30]. Caylakli et al. determined that maxillary sinus mucoceles make up 10% of all mucoceles [24].

**Fronto-ethmoid Mucoceles**

Mucoceles arising from the frontal sinus present with a variety of clinical signs such as decreased visual acuity, visual field abnormalities, proptosis, ptosis, periorbital swelling, displacement of the globe, restricted ocular movements and choroidal folds. The patient’s visual activity is usually not affected unless compression of the optic nerve occurs from the mass effect of the mucocele. Meningitis, meningoencephalitis, pneumocephalus, brain abscess, seizures, and cerebrospinal fluid fistulas are other serious sequelae. The most common presenting symptom was proptosis in 61.5% of the patients reviewed. Forehead swelling and nasal congestion were each present in 38.5% of patients. Other presenting symptoms include orbital swelling, forehead sinus cutaneous fistula, orbital cellulitis, and swelling of medial canthus [25]. Possible initial sequelae include infection, cerebrospinal fluid fistula, meningitis, thrombosis, abscess, paresthesia, and headache. Less common sequelae include cosmetic abnormality, pyomucocele, intracranial abscess, headaches, and osteomyelitis. Orbital involvement is rare, but includes vision loss, diplopia, superior ophthalmic vein thrombosis, cavernous sinus thrombosis, and proptosis. Sinus outflow obstruction from trauma, inflammation, previous surgery, tumors, and polyps are exacerbating factors [6]. The definitive treatment of frontal mucoceles is surgical [32]. Until the 1980’s, mucoceles were exenterated mainly using external approaches. Leading surgical techniques were Lynch–Howarth [33] incision and osteoplastic frontal sinusotomy as described by Bockhmul et al. [34] and by Weber et al. [35]. Traditionally, non-endoscopic treatment of FOE mucoceles has involved trephination procedures (Figure III) or osteoplastic flap with mucosal extirpation and fat obliteration.

In 1989, Kennedy et al. proposed an alternative endoscopic approach that showed no recurrences after a mean followup of 18 months [36]. Today, marsupialization of mucoceles using endoscopic sinus surgery is considered the first choice of treatment [19,30,32]. It is a safe method with both low morbidity and recurrence rates (Figure IV). External approaches to frontal sinus are still used, alone or combined with ESS, but no clear-cut definition of the limits of a successful endoscopic procedure has been given [32,37,38].

**Figure II.** The coronal plane CT image showing a mucocele (asteriks) in the right frontal sinus base that thinned the bone structure (A). In addition, thinning (arrows) and deformation can be observed in the right sinus roof (B) and anterior wall of the left frontal sinus (C).

**Figure III.** Frontal sinus trephination in a patient that cannot be entubed due to giant hemangioma of the tongue.

**Figure IV.** Endoscopic marsupialization of fronto-ethmoid mucocele.

A survey focusing on the current management of frontal sinus disease in the United Kingdom showed that 54% of surgeons are using an external approach to treat frontalsinus mucoceles independent of their location; only 39% attempt an endoscopic drainage if the cyst lies sufficiently close to the midline [39]. Correa 21 stated that patients with a narrow anterior-posterior diameter of the frontal recess, a highly comparmentalized...
frontal sinus, an extensive polypoid degeneration of the frontal sinus mucosa, or those with high thickened secretions are candidates for obliteration [40]. Successful frontal sinus obliteration requires meticulous removal of all visible sinus mucosa and the inner cortex of the sinus wall. A wide variety of implant materials for obliteration has been described. Autologous fat transplantation is regarded as the gold standard [41]. The most common reasons for choosing an osteoplastic flap procedure were erosion of the anterior frontal sinus wall requiring forehead reconstruction, far lateral located lesions, or unfavorable frontal recess anatomy with extensive scarring [25]. We choose external approaches in far lateral located lesions, anterior frontal sinus wall’s defects, and huge lesions expanding into the intracranial region (Figures V, VI). On the other hand, M. Trimarchi treated frontal sinus mucoceles with lateral extension by ESS Surgical technique included anterior ethmoidectomy and either Type I, Ila or Iib frontal sinus drainage according to Draf [42,43]. Eloy JA reported the use of a balloon dilation method for drainage of a large frontal sinus mucocele (44).

**Maxillary Sinus Mucoceles**

Maxillary sinus mucoceles are more prevalent in Japan, where they are usually reported as following Caldwell-Luc maxillary sinusectomy [24,45,46]. Mucoceles that develop following Caldwell-Luc operations are presumed to form as a result of entrapped sinus mucosa. Although one of the theories about development of mucocele is chronic infection, Busaba et al. compared the bacteriology of maxillary sinus mucoceles to chronic sinusitis and reported that the data does not support infection as the main origin of nontraumatic maxillary sinus mucocele [47]. Caylakli et al. state that in 64% of a series of 14 patients, the etiological cause remains uncertain. The symptoms of mucoceles are related to their expansion and subsequent pressure and the obstruction of surrounding anatomic structures (Figure VII). Antral mucoceles are commonly reported to present as a painless bulging of the cheek. Medial expansion of the wall of the maxillary sinus into the nasal cavity displaces the inferior turbinate and causes the nasal obstruction [48]. Superior expansion of the antrum into the inferior orbit can cause displacement of the orbital contents and visual changes. Downward displacement into the area of the alveolus can even cause a loosening of teeth [46].

Retention cysts are common in the maxillary sinus and may be found on imaging studies in approximately 9% of the population. They are thought to form due to obstruction of the ducts of seromucous glands in the sinus lining, which results in an epithelium-lined cyst containing mucous or serous fluid. They develop under mucous membranes of the sinus which explains why they are so thin-walled. Radiographically, the cyst is a rounded, dome-shaped, soft tissue mass, most commonly situated on the floor of the maxillary sinus, often containing clear, yellowish fluid. Mucoceles are associated with obstruction of the duct or natural ostium of any of the paranasal sinuses and grow under the peristome. Periosteum contributes to construction of the cystic wall, and, as a result, the wall of mucocele becomes thick and tough. The growing site of the mucocele is under the periosteum, whereas retention cysts grow under the mucosa of the sinus. This explains why retention cysts are non-expanding, well

Figure V. A huge fronto-ethmoidal located mucocele (asterisk) is shown. In the coronal CT (A) and T1 (B) and T2-weighted MR images (C, D) the expansile mucocele (arrows) causes thinning (arrowheads) of the frontal sinus walls.

Figure VI. The coronal T1-weighted MR image after surgery (E). Air (asterisk) is observed in the left fronto-ethmoidal cavity.

Figure VII. In the photograph, expansion of the maxillary bone and in CT scan; medial expansion of the wall of the maxillary sinus is seen.
circumscribed, mucosa-covered masses, whereas mucoceles exhibit an osteolytic capacity with a tendency to expand along the path of least resistance [24,48,49].

In the treatment of maxillary mucoceles, surgical techniques including external approaches, marsupialization, Caldwell-Luc procedures, and endoscopy are used [45,46,49]. There was no relapse in a 14-patient series by Caylakli et al. in which endoscopic ethmoidectomy and middle meatal antrostomy were applied. We consider that an extensive marsupialization of the mucocele using middle meatal antrostomy is sufficient for treatment (Figure VIII).

Figure VIII. Endoscopic marsupialization of the maxillary sinus mucocele.

Sphenoid Sinus Mucoceles
Sphenoid sinus mucocele is a rare condition which represents only 1–2% of all paranasal sinus mucoceles [31]. Three pathogenetic mechanisms have been postulated for sphenoid sinus mucocele: submucosal oedema, secretory duct obstruction, and ostial obstruction. Submucosal oedema may result from allergic rhinitis causing ostia narrowing with obstruction; ostial obstruction may also be due to chronic sinusitis, polyposis, or both [50]. Sphenoid sinus mucocele formation has been reported as a possible late complication of radiotherapy to the head and neck [51]. Radiation causes scarred mucosa which may occlude the sinus ostium.

Presenting symptoms are non-specific, and patients may be asymptomatic. The clinical symptoms and signs of sphenoid sinus mucocele are directly related to the sinus anatomy and its contiguous neurological and vascular structures. Important structures are described which are contiguous with the sphenoid sinus: cranial nerves II to VI, cavernous sinus, carotid artery, sphenopalatine artery and nerve, pterygoid canal and nerve, dura, and pituitary gland. Patients with sphenoid sinus mucocele become symptomatic when any of these structures are displaced or compressed; mucocele commonly spreads along the path of least resistance, anteriorly, into the nasopharynx, or anterolaterally, toward the orbit. Sphenoid sinus mucocele is also more likely to present with visual changes than with inflammatory disease. Visual loss in sphenoid disease has been attributed to optic neuritis or compression [52]. Hejazi et al. reviewed 130 cases of symptomatic sphenoid sinus mucoceles reported up to 2001. The patients presented with these symptoms, in order of decreasing frequency: headache (87%), ocular disorders (85%), amaurosis (58%), oculomotor palsy (55%), nasal symptoms (38%), endocrine disorders (4 of 130) and anhypo-pituitarism (1 of 130) [53]. Intracranial extension is unusual in sphenoid sinus mucocele. If extension occurs, it is commonly into the suprasellar region or anterior cranial fossa. Computerized tomography provides valuable information regarding bony erosion, while MRI enables more precise visualisation of the involvement of the optic nerve, dura, pituitary gland, cavernous sinus, and carotid artery [52].

Mucoceles may show various imaging features, depending on their contents. They often have low attenuation on CT, a low signal on T1-weighted MRI, and a high signal on T2-weighted MRI, due to their high water content. There is usually no enhancement, or at most marginal enhancement, on CT and T1-weighted MRI, whereas many of the lesions in the differential diagnosis show contrast enhancement. Thus, the diagnosis of mucocele can be suspected pre-operatively and differentiated from craniopharyngioma, pituitary tumor, dermoid, chordoma, epidermoid, aneurysm, metastatic lesions and, in rare cases, an arachnoid cyst. In mucoceles with a high protein content, T1-weighted MRI shows a lesion which is homogeneously hyperintense centrally with a surrounding rim of hypointense mucosa [53]. Endoscopic drainage is safe and effective in eradication of disease.

Middle Concha Bullosa Mucoceles
A concha bullosa alone is usually asymptomatic. An enlarged concha bullosa may obstruct the middle meatus or infundibulum. Headaches, orbital pain, exophthalmia, nasal discharge, postnasal drip, nasal obstruction, and hyposmia are all possible symptoms of a concha bullosa mucopyocele. A concha bullosa mucopyocele may also manifest as migraine headaches [54].

Competing interests
The authors declare that they have no competing interests.

References