Nasopharyngeal Carcinoma with Peri-Eustachian Tube Extension: A Rare Pathway of Spread

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Keywords: Nasopharyngeal carcinoma; lymph node; MRI.

Abstract
Nasopharyngeal carcinoma (NPC) is a widespread malignant disease in endemic regions such as South Asia. Palpable nodal metastasis in the neck or involvement of adjacent structures and ear complaints are common presentations.

A 53 years old male presented with six months history of progressive unilateral face pain and a history of ear discharge for one month. Ultrasound examination of the neck showed multiple bilaterally enlarged lymph nodes. Magnetic resonance imaging (MRI) scan depicted heterogeneous mass in the right side of the nasopharynx. Nasopharyngeal histopathology revealed it to be a non-keratinizing squamous cell carcinoma. The patient underwent radiotherapy for disease management. MRI helps detect disease in the early stage as well as for precise extension of the disease.

Introduction
Nasopharyngeal carcinoma (NPC) is the most common primary malignancy arising from the nasopharynx’s epithelium. It is commonly diagnosed in between 40 and 60 years. Females are less commonly affected than males. NPC is a classic disease with clinical behaviour, epidemiology, and histopathology different from squamous cell carcinomas in the head and neck [1].

In advanced disease, palpable neck mass and ear complaints are common in nasopharyngeal carcinoma (NPC). The vast majority of data propose Epstein–Barr virus (EBV) as the predominant pathogenic factor in NPC [2].

This case report discusses a patient with NPC presented with extensive skull base disease, lymph nodes involvement, and ear extension.

Case presentation
A previously healthy 53-year-old man presented to our hospital with ear discharge and pain on the right side of the face that started six months before this visit. There were associated hearing impairment and fullness in the bilateral ear, but no balance disorders. Nasal blockage and postnasal drip were experienced sometimes, but there were no nasal discharge and epistaxis. There was no history of injury or any neurological deficit. Physical examination of the patient revealed palpable lymph nodes in bilateral neck levels. He had a 24 pack/year smoking history but was otherwise absent known cancer risk factors.

Investigations
The patient underwent an ultrasound of the neck, which showed “numerous round to oval hypoechoic nodes”. The larg-
lymph nodes seen at various levels on both sides of the neck. There were also multiple heterogeneously enhancing enlarged changes of sinusitis. Nasal turbinates and meati were normal.

The bilateral maxillary, frontal and ethmoid sinus, representing along the tube contrast enhancement in its entire length s/o tumour extension chain tube blockage. The right eustachian tube also showed post-

bilateral posterior choanae. Fluid signal intensities were seen in

erogenous parotid lymph node on the right side cranial fossa, clivus, involves the right cavernous sinus with hetero
geneous lesion described above shows post-contrast heterogeneous en

rangement Imaging (MRI) of skull base that included cavernous si

(fossa through foramens and causing expansion of the right cavern

us) (Figure 2). Mucosal thickening was present in the bilateral maxillary, frontal and ethmoid sinus, representing changes of sinusitis. Nasal turbinates and meati were normal. There were also multiple heterogeneously enhancing enlarged lymph nodes seen at various levels on both sides of the neck.

Following oncological consultation, a biopsy of a lymph node was performed. Histological examination revealed the presence of “lymphatic metastasis of undifferentiated cell carcinoma with Regaud growth pattern, likely nasopharyngeal neoplasm”. In addition, Rhinoscopy confirmed an "undifferentiated nasopharyngeal carcinoma", suggesting a stage IV (T4 N2cM0; AJCC 8th Edition) nasopharyngeal carcinoma.

Treatment

Once the diagnosis was obtained, the patient was referred to an oncology department. There, he was treated for six weeks with radiotherapy, cisplatin and corticosteroids.

Outcome and follow-up

After six weeks patient was came to department for follow-up with relief of ear complains.

Discussion include a very brief review of similar published cases

NPCs are one of the head and neck malignancies common in the endemic geographical regions such as the southern region of Asia, including Bangladesh, Bhutan, India, Pakistan, Nepal, and Sri Lanka [1].

Consumption of salted fish has been related to the carcinogenic compound. Nitrosamine is one of the most frequently mentioned etiologic factors. The EBV genome is commonly detected in the biopsy specimens of NPC. Hence it has been considered to play an oncogenic role in this tumour[2]. However, in this particular patient, there was no history of exposure to carcinogens.

As for EBV infection, while it is widespread and affect over 90% of the Asian adult population, it is usually asymptomatic. The NPCs associated with EBV are of undifferentiated form [3].

Symptoms of NPC presentation can be placed in four groups. They are 1. Nasal symptoms such as discharge, nasal obstruction and epistaxis (these symptoms are related to the presence of tumour mass in the nasopharynx); 2. Otologic symptoms include tinnitus and deafness (related to the Eustachian tube dysfunction caused by the anteroposterior involvement of the tumour to the paranasopharyngeal space) ; 3. Cranial nerve palsies, commonly fifth and sixth cranial nerves (associated with the superior extension of the tumour leading to skull base erosion), might experience facial pain, headache, diplopia, and numbness; and 4. Neck masses usually appears in the upper 1/3rdneck [4].

Patients with an early stage of the NPC may present with complaints related to the fullness of the ear.
Eustachian tube dysfunction may be caused by a small tumour in the fossa of Rosenmüller of the nasopharynx. The tumour usually expanded outward through the mucosa and sub-mucosa, along with the muscle bundles within the neuro-vascular planes or the fibro-fatty tissue planes. The Eustachian tube and middle ear or auditory nerve involvement with the resulting correlated aural manifestations also happen[5].

Early-stage NPC is challenging to diagnose clinically because of its hidden location in the nasopharynx [6]. Moreover, the presentations of NPC are variable. Still, they may also remain symptomless for a long time, making it harder to disclose the occult site of presentation. Moreover, sometimes patients consult doctors of different specialties who have little experience in managing nasopharyngeal carcinoma [7].

In this NPC case, the patient had a month of progressive ear discharge and six-months of the right side face pain. There was an associated feeling of fullness and hearing impairment in the bilateral ear. A thorough nasoendoscopic examination can make it possible to detect an early-stage NPC.

WHO classified NPCs in 2005 into three main types. They are keratinizing squamous cell carcinoma, non-keratinizing squamous cell carcinoma, and basaloid squamous cell carcinoma. Histopathology demonstrates, in this case, revealed that the tumour was a non-keratinizing squamous cell carcinoma. These classifications of tumours are commonly recognized as radiosensitive, which differentiate these from squamous cell carcinoma and have a stronger association with EBV [8].

MRI is advised to be an accurate examination to detect NPC. MRI can diagnose subclinical NPCs that are hidden at endoscopy. NPCs in MRI typically present low signal intensity on T1-weighted images and intermediate signal intensity on T2-weighted images. Most NPCs arise from the posterolateral recess of the pharyngeal wall (Rosenmüller fossa) (82%), midline (12%) and 6-10% of patients showed normal nasopharyngeal mucosa appearance at endoscopy[9]. MRI scans reveal soft tissue thickening in the nasopharynx and involvement of adjacent structures. No further examinations are needed.

Significant prognostic factors for survival in NPC are the stage, WHO type, and age at diagnosis[10]. Therefore, this particular patient is expected to have a good prognosis.

Learning points/take home messages 3-5 bullet points

- In this case, effacement of the fat in torus tubarius and the fossa of Rosenmüller region with asymmetrical fullness is seen in early nasopharyngeal carcinomas.
- Subtle effacement with fullness along the right lateral nasopharyngeal wall, along with pathological right parotid lymph node, and bilateral mastoid effusion, due to Eustachian tube dysfunction in this patient with biopsy-proven nasopharyngeal carcinoma.
- The most common imaging appearance of NPC is a mass centered at the lateral nasopharyngeal recess with deeper extension (adjacent spaces) and associated cervical nodal disease.

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