



NASOPHARYNGEAL CARCINOMA 鼻咽癌之評估與處置

高雄長庚醫院耳鼻喉部

NASOPHARYNGEAL CARCINOMA 鼻咽癌之評估與處置

金宇彬經紀公司證實

罹患“鼻咽癌”治療中

高雄長庚醫院耳鼻喉部

Division of Pharynx

- The nasal part –

NASOPHARYNX/ EPIPHARYNX

(extends from base of skull to soft palate)

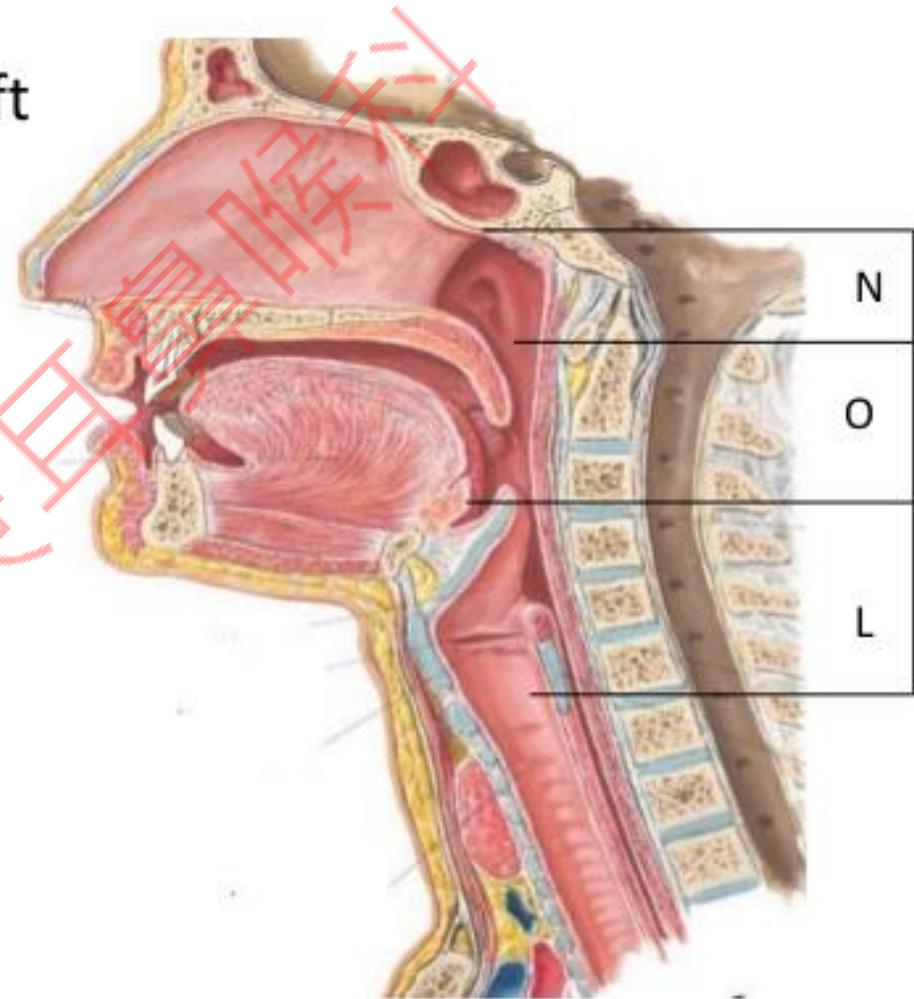
- The oral part – **OROPHARYNX**

(extends from hard palate to hyoid bone)

- The laryngeal part –

**LARYNGOPHARYNX/
HYPOPHARYNX**

(extends from upper border of epiglottis to lower border of cricoid cartilage)



Boundaries

- anteriorly: posterior nares and posterior margin of nasal septum
- inferiorly: soft palate
- superiorly: basisphenoid and basiocciput
 - roof of the nasopharynx is called the vault (or fornix) of the pharynx, where the mucosa firmly attaches to the sphenoid and pharyngobasilar fascia
- posteriorly: C1 and C2
- laterally
 - the pharyngeal opening of the Eustachian tube is located in the centre of the lateral wall
 - lymphoid tissue aggregates, also known as the tubal tonsil occur around the opening of the Eustachian tube
 - the Fossa of Rossmüller lies between the posterior margin of the Eustachian tube and the posterior wall of the nasopharynx

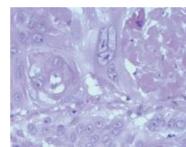
Incidence and etiology

- NPC的癌症型態為SCC，最常出現在Rosenmuller fossa
- 歐美盛行率:< 1 / 10萬
- 阿拉斯加及中國南部尤其廣東省較盛行,香港男(20-30/ 10萬)及香港女(15-20 / 10萬)
- 中國人移民到北美依舊有較高盛行率，但是在北美出生的第二代則否,病因決定於基因,民族及環境
- 跟NPC有關的因素有salted fish、EBV、multiple chromosome(14q,16p,1p,12q,4q)。有NPC家族史的人得NPC是平常人的六倍

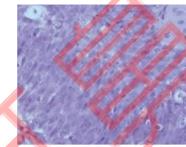
Histopathology

- WHO(1978) histologic classification of NPC

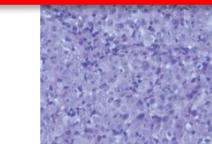
- Type I: typical keratinizing SCC
- Type II: nonkeratinizing epidermoid carcinomas
- Type III: undifferentiated or poorly differentiated carcinomas



type I



type II



type III(最多)

- | Region | Percentage |
|---|------------|
| 北美 | 25% |
| 中國南部 | 3% |
| 切片常切出mixed pattern, 所以WHO新分法: 組織學上只先分: | |
| (1)SCC | |
| (2)nonkeratinizing carcinomas (再分Differentiated/ undifferentiated carcinomas) | |
| • 使用radiotherapy治療undifferentiated carcinomas的local tumor control rate比較高 | |
| • undifferentiated carcinomas的 distant metastasis incidence比較高。 | |

Clinical presentation

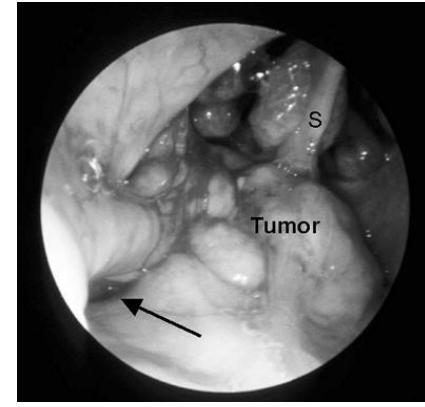
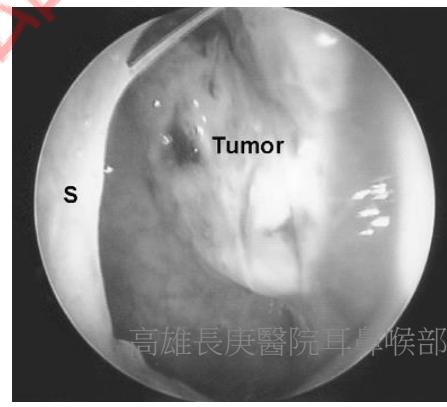
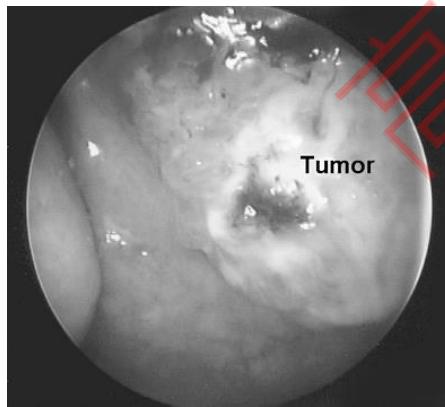
- History
 - Cranial nerve involvement: headache, diplopia, facial numbness
 - Tumor infiltrates skull base→ headache
 - Tumor extends to cavernous sinus, the 3rd, 4th, and 6th cranial nerves→ diplopia
 - Tumor extends to foramen ovale (CN5)→ facial pain and numbness
 - Cervical node metastasis: mass in the neck
 - upper neck painless mass (最常見; 容易雙側)
 - Clinical triad: neck mass, nasal obstruction with epistaxis, serous otitis media
 - 越下方的anatomy比率越高:**neck mass (76%) > nasal symptom(73%)>ear symptom(62%)>CNS symptom (只有20%)**
 - Distant metastasis: vertebra, liver, and lung

Diagnosis

- History and physical examination
- Image studies: nasopharyngoscopy, chest radiograph, CT/MRI of the nasopharynx, skull base, and neck
- Serology

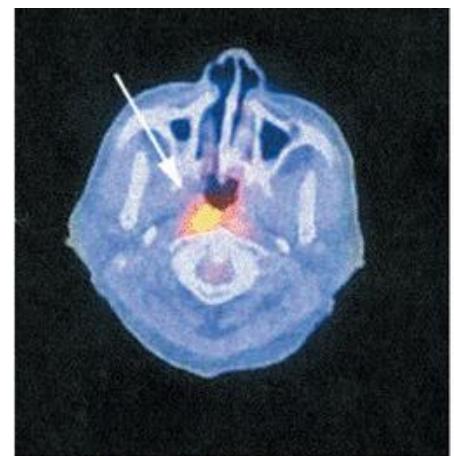
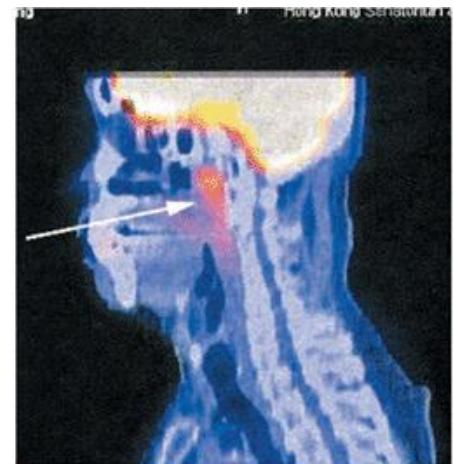
Endoscopic Examination

- 要確診NPC還是需要positive biopsy
- rigid Hopkin telescopes: 0° and 30°
- 在NSD的病人可以用70° 看對側或是從soft palate下方看
- Flexible endoscope就算只塞進一個鼻孔也可以檢查整個nasopharynx



Imaging Studies

- 多用CT或MRI, PET對於recurrent或是persistent NPC比前兩者好
- CT看bone erosion(skull base), perineural spread(從foramen ovale入顱內, 可偵測involve cavernous sinus但無顱底骨侵犯), bone regeneration after therapy (代表腫瘤根除)
- MRI :可分別tumor跟inflammation, retropharyngeal/ deep cervical nodal metastases, bone marrow infiltration(增加遠端轉移的機會)
- Bone scans, liver scintigraphy, and marrow biopsy



Diagnosis: Serology

- EBV: infectious mononucleosis, Burkitt lymphoma, NPC
- EBV-specific antigens 分三個group
 - early replicative antigens : IgA anti-EA (early antigen)
 - latent phase antigens
 - late antigens: IgA anti-VCA (viral capsid antigen):
比較敏感但特異性低，若升高則subclinical NPC機會約3%~5%，一年內診斷NPC是一般人30倍，它跟disease的stage有關，但作為tumor marker去評估是否根除或復發則未有定論
- anti-EBV titer升高,NPC的機會是一般人30倍
- anti-EBV ,IgA anti-EA, IgA anti-VCA被認為有診斷價值 (EA,VCA->IgA)
- **EBNA (EBV-associated nuclear antigen)的敏感跟特異性都超過92%**
- EBNA及IgA anti-EA特異性較高 (EBNA,EA 特異性高)
- cell-free DNA of EBV 有用為NPC tumor marker
- 血清學的診斷僅能用以輔助組織學的診斷，而可在NPC病人電療後的追蹤上有幫助

Treatment

- NPC是radiosensitive的，所以多用RT治療。
- 比較advance diseases會使用 concurrent chemoradiotherapy(CCRT)
- persistent或recurrent disease:
 - 在neck:以radical neck dissection做為 salvage surgery 比再接受一次radiotherapy的效果好
 - 在nasopharynx:可使用stereotactic radiotherapy、brachytherapy(定義是 radiation source直接放到tumor 裡面，最常使用的是金198Au)、nasopharyngectomy、external radiotherapy + concurrent chemotherapy
- 如果病人有distant metastasis, standard treatment是 cisplatin + 5-FU。

Treatment : *Radiotherapy*

- 成功率:
- T1 and T2 tumors : 75% to 90%
- T3 and T4 tumors : 50% to 75%
- N0 and N1: 90%
- N2 and N3 cases: 70%
- 3D conformal radiotherapy and IMRT:挑戰在於決定腫瘤的邊界
 - 當腫瘤接近 dose-limiting organs
 - 精準的控制 primary tumor 及 neck 交界處及 upper neck 的劑量
 - 極佳的 locoregional control , 兩年後唾液腺功能可以回復
 - 對 recurrent NPC, short-term control 效果不錯

Chemotherapy

- 在 advanced locoregional disease 會使用 CCRT
- cisplatin 在 RT 前中後都可以使用
 - neoadjuvant, concurrent, and adjuvant chemotherapy with radiotherapy
- 1997 study 發現 CCRT 比單獨 RT 好 (同時作最好)
- Neoadjuvant studies: relapse-free survival 改善但 overall survival 沒改善
- adjuvant chemotherapy: relapse-free survival, overall survival 都沒改善.