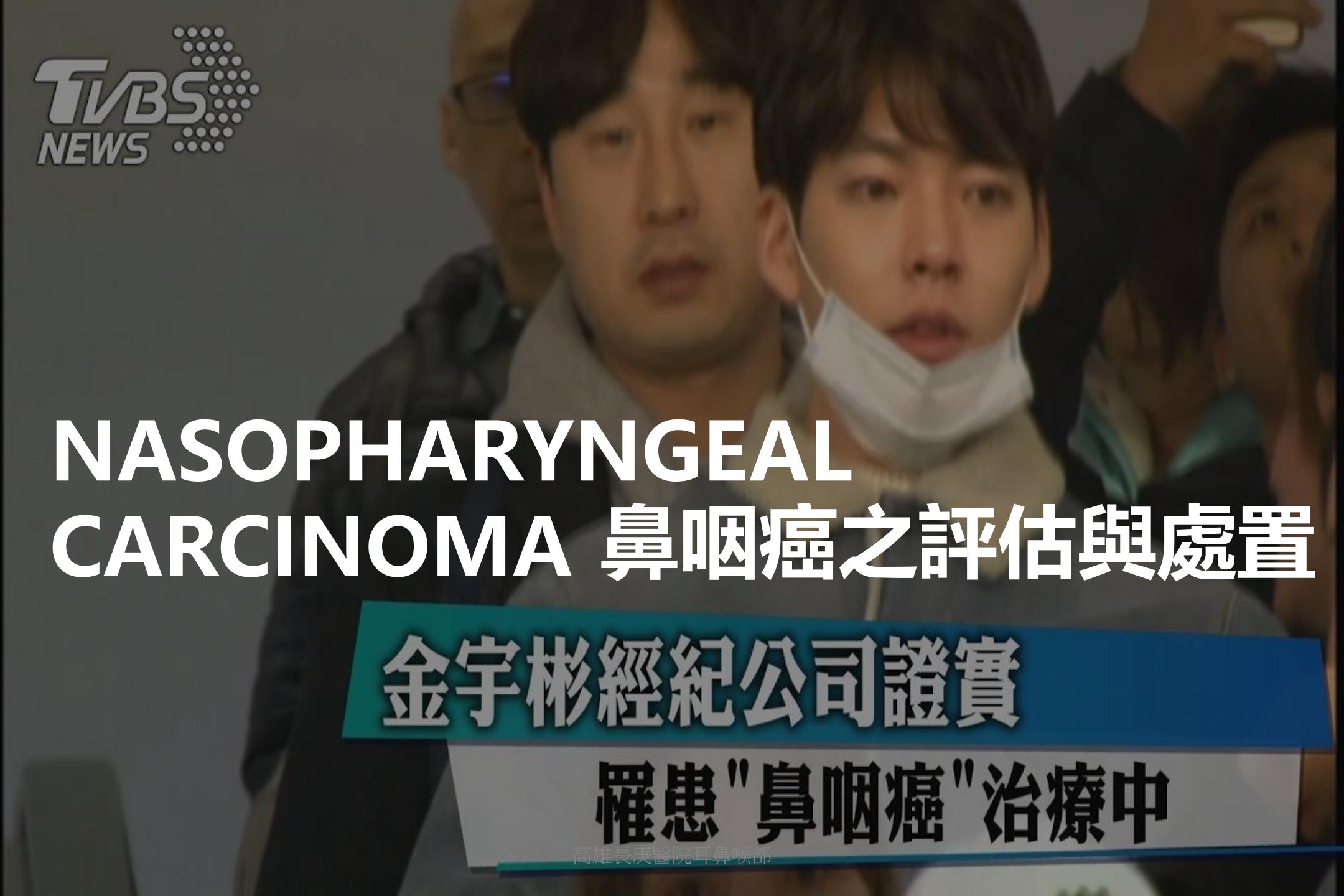




# NASOPHARYNGEAL CARCINOMA 鼻咽癌之評估與處置

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

羅患“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)

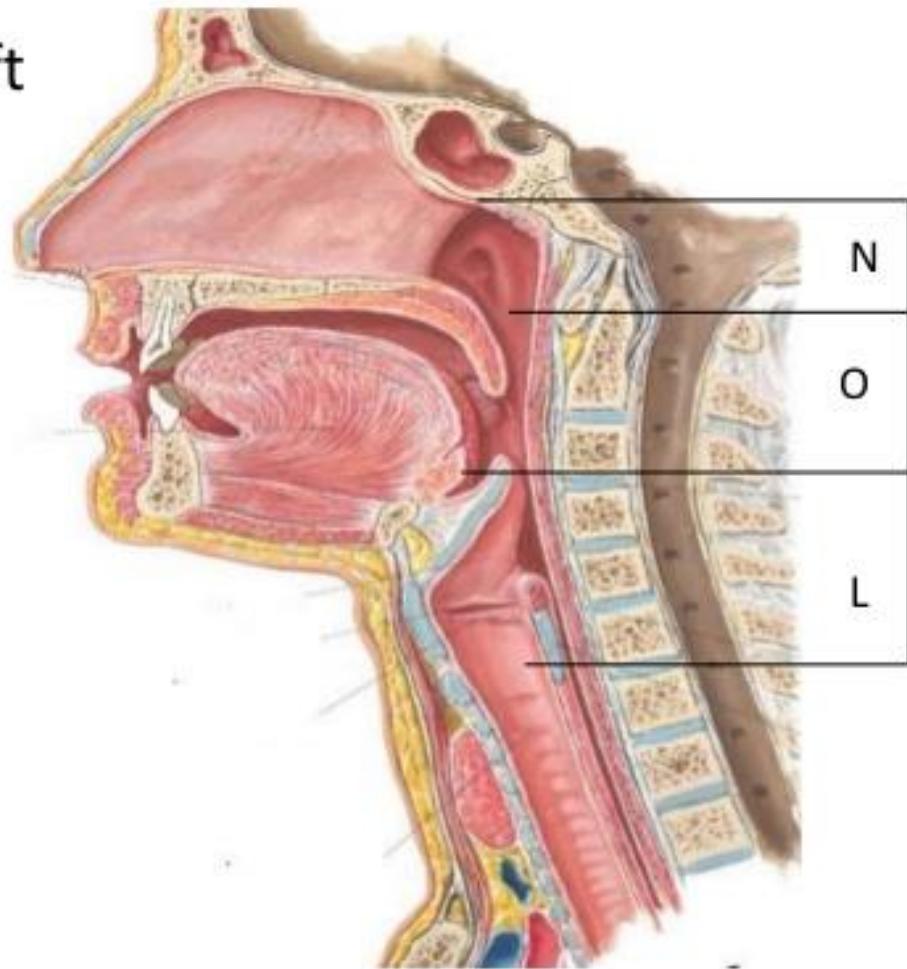
- The oral part – **OROPHARYNX**

(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

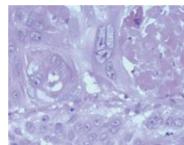
- anteriorly: posterior nares and posterior margin of nasal septum
- inferiorly: soft palate
- superiorly: basisphenoid and basiocciput
  - roof of the nasopharynx is called the vault (or fornix) of the pharynx, where the mucosa firmly attaches to the sphenoid and pharyngobasilar fascia
- posteriorly: 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 Rossenmüller lies between the posterior margin of the Eustachian tube and the posterior wall of the nasopharynx

# Incidence and etiology

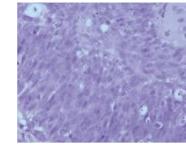
- NPC的癌症型態為SCC，最常出現在Rosenmuller fossa
- 歐美盛行率:< 1 / 10萬
- 阿拉斯加及中國南部尤其廣東省較盛行,香港男(20-30/ 10萬 )及香港女(15-20 / 10萬 )
- 中國人移民到北美依舊有較高盛行率，但是在北美出生的第二代則否,病因決定於基因,民族及環境
- 跟NPC有關的因素有salted fish、EBV、multiple chromosome(14q,16p,1p,12q,4q)。有NPC家族史的人得NPC是平常人的六倍

# Histopathology

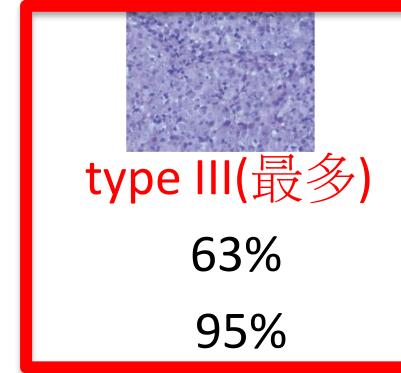
- WHO(1978) histologic classification of NPC
  - Type I: typical keratinizing SCC
  - Type II: nonkeratinizing epidermoid carcinomas
  - Type III: undifferentiated or poorly differentiated carcinomas



type I



type II



type III(最多)

- 北美 25% 12%
- 中國南部 3% 2%

63%

95%

- 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

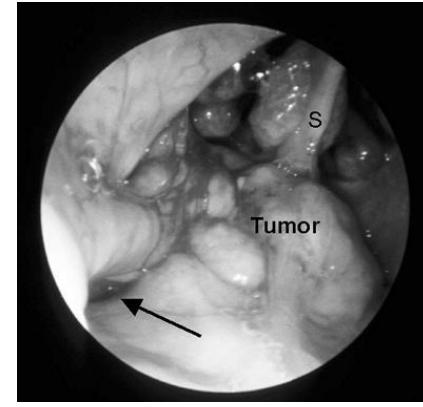
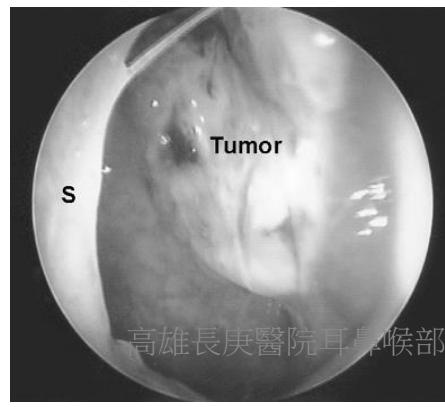
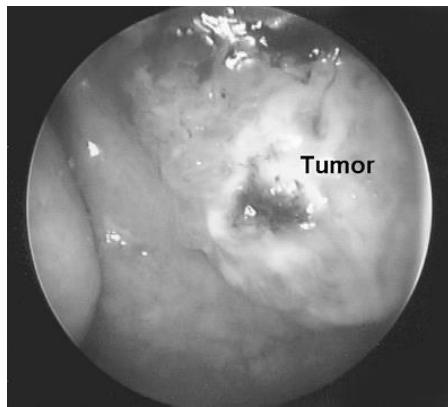
- History
  - Cranial nerve involvement: headache, diplopia, facial numbness
    - Tumor infiltrates skull base→ headache
    - Tumor extends to cavernous sinus, the 3rd, 4th, and 6th cranial nerves→ diplopia
    - Tumor extends to foramen ovale (CN5)→ facial pain and numbness
  - Cervical node metastasis: 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

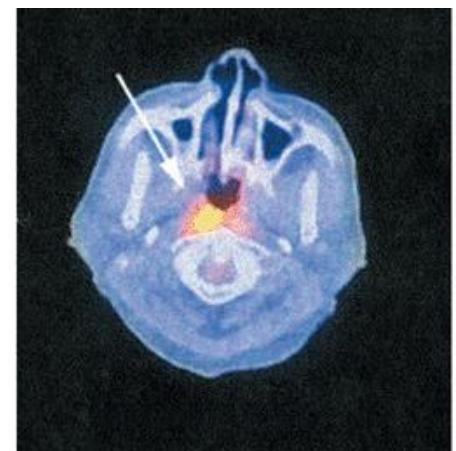
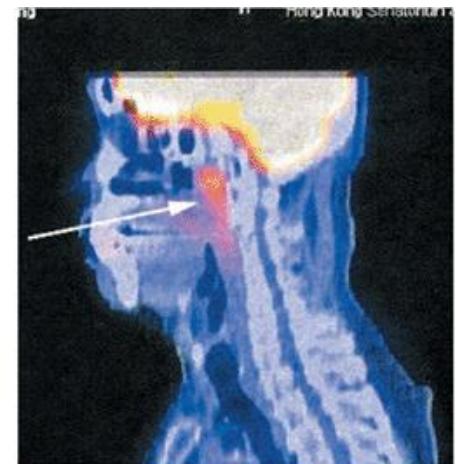
# Endoscopic Examination

- 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.