

Minimally Invasive Endoscopic Skull Base Surgery for Pituitary Tumors

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Pituitary Lesions

- ***Pituitary adenomas***
 - Non-Secretory / Non-Functional
 - Secretary / Functional
 - PRL-Secreting / Prolactinoma
 - ACTH-Secreting / Cushing's Disease
 - GH-Secreting / Acromegaly
 - Apoplexy
- ***Rathke's cleft cysts***
- **Craniopharyngiomas**
- Meningiomas
- Rarities: pituicytoma, hypophysitis, clival plasmacytoma, germ cell tumors
- *Work-force of endoscopic skull base surgery*

Pituitary Adenomas

- 10-15% of all primary intracranial neoplasms (third most common primary intracranial tumor)
- 3rd – 6th decade of life
- “Benign”.
- Presentation:
 - Incidental
 - Mass effect:
 - Optic chiasm -> bitemporal vision loss
 - Pituitary -> pituitary insufficiency.
 - Hypersecretory states

2017 WHO Pituitary Classification

- Classification of pituitary tumors according to cell lineage/transcription factors.

Pituitary adenomas	Somatotroph adenoma Lactotroph adenoma Thyrotroph adenoma Corticotroph adenoma Gonadotroph adenoma Null-cell adenoma Plurihormonal and double adenomas
Pituitary carcinoma Pituitary blastoma Tumors of the posterior pituitary	Pituicytoma Granular cell tumor of the sella Spindle cell oncocytoma Sellar ependymoma Gangliocytoma and mixed gangliocytoma-adenoma Neurocytoma Paraganglioma Neuroblastoma
Neuronal and paraneuronal tumors	Adamantinomatous craniopharyngioma Papillary craniopharyngioma Meningioma Schwannoma Chordoma, NOS Chondroid chordoma “Dedifferentiated” chordoma Solitary fibrous tumor/hemangiopericytoma Grade 1 SFT/HPC Grade 2 SFT/HPC Grade 3 SFT/HPC
Craniopharyngioma	
Mesenchymal and stromal tumors	
Hematolymphoid tumors Germ cell tumors	Germinoma Yolk sac tumor Embryonal carcinoma Choriocarcinoma Teratoma, NOS Mature teratoma Immature teratoma Teratoma with malignant transformation Mixed germ cell tumor
Secondary tumors	

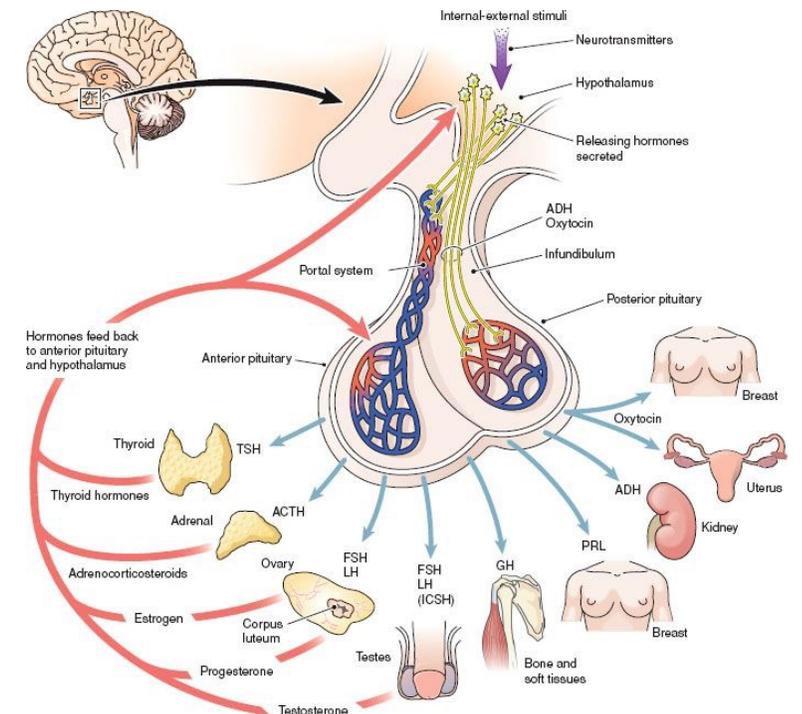
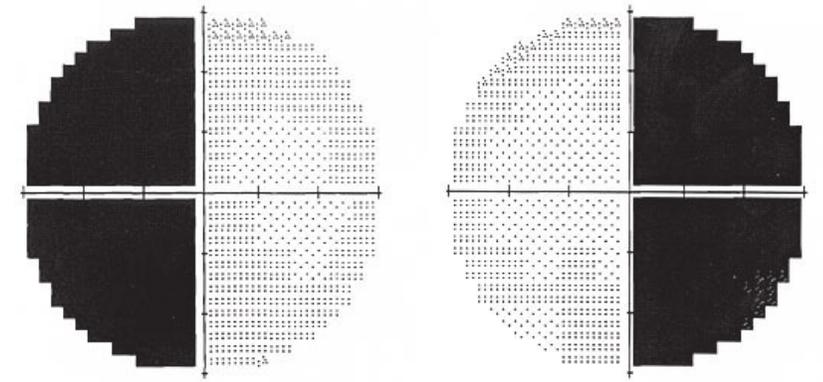
Table 2 Adenohypophyseal cell lineage basis for the 2017 classification of pituitary adenomas

Lineage	Main transcription factors and other co-factors	Adenohypophyseal cell
Acidophilic lineage	PIT-1	Somatotrophs
	PIT-1, ER α	Lactotrophs
	PIT-1, GATA-2	Thyrotrophs
Corticotroph lineage	T-PIT	Corticotrophs
Gonadotroph lineage	SF-1; GATA-2, ER α	Gonadotrophs

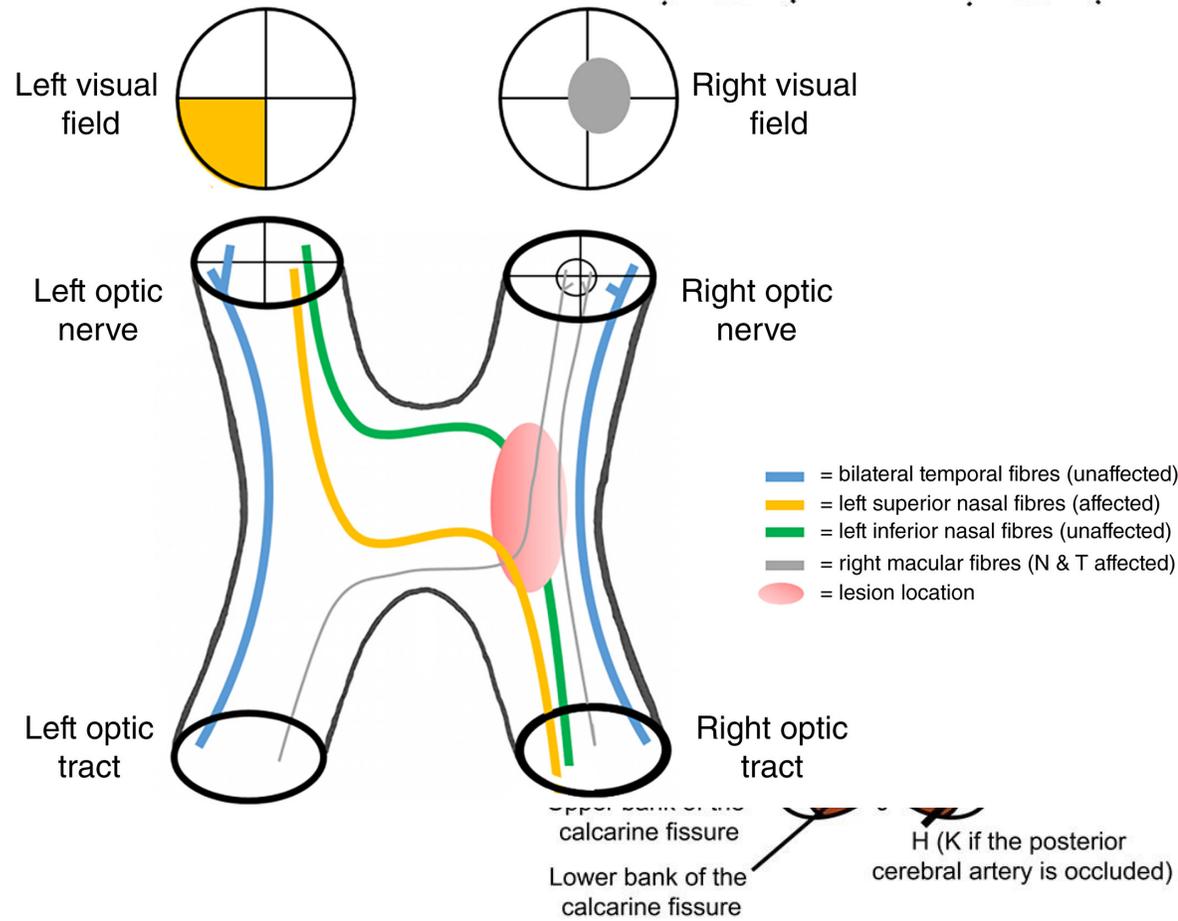
PIT-1 pituitary-specific POU-class homeodomain transcription factor 1, *ER α* estrogen receptor α , *GATA-2* member of the GATA family of zinc-finger transcriptional regulatory proteins, *T-PIT* T-box family member TBX19, *SF-1* steroidogenic factor 1 [48; for review]

Pituitary Adenoma Work-Up

- Imaging: T1 post contrast coronal and sagittal planes
- Vision exam; visual fields test (Humphrey)
- Endocrine work-up
 - Diagnose a hyper-secretory pituitary tumor
 - Establish baseline
 - TSH, fT4, PRL, ACTH, 7am cortisol, FSH/LH, free/total testosterone, estrogen, GH, IGF-1
 - BMP to check Na
- Advanced endocrinological tests
 - Cushing's disease work-up



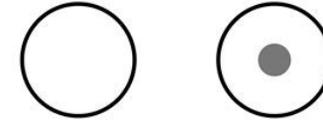
Visual Field Defects



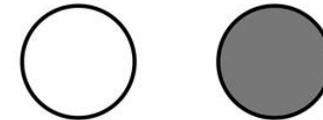
t eye

Visual field deficit

A) Central scotoma



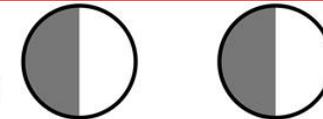
B) Monocular vision loss



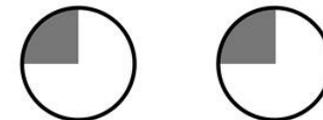
C) Bitemporal hemianopia



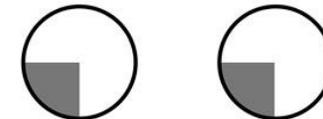
D, G, & H) Contralateral homonymous hemianopia



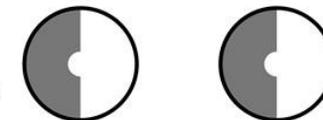
E & J) Contralateral superior quadrantanopia



F & I) Contralateral inferior quadrantanopia



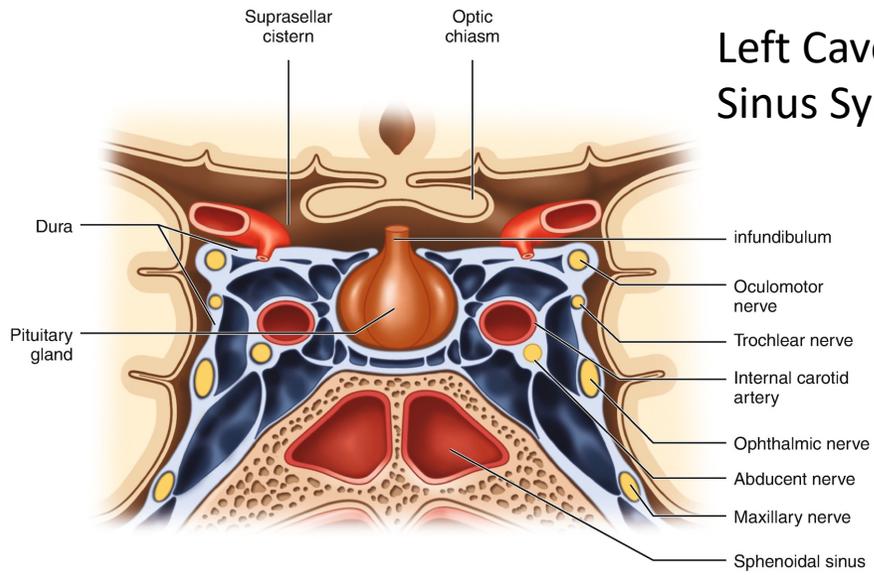
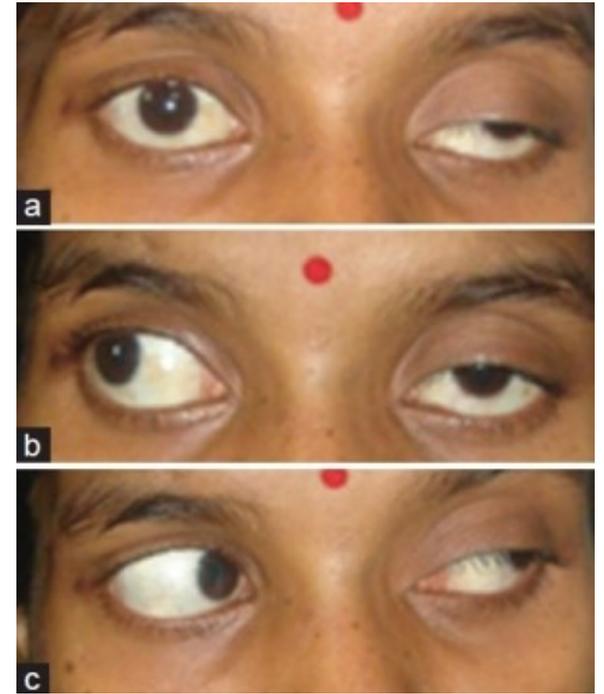
K) Contralateral homonymous hemianopia with macular sparing



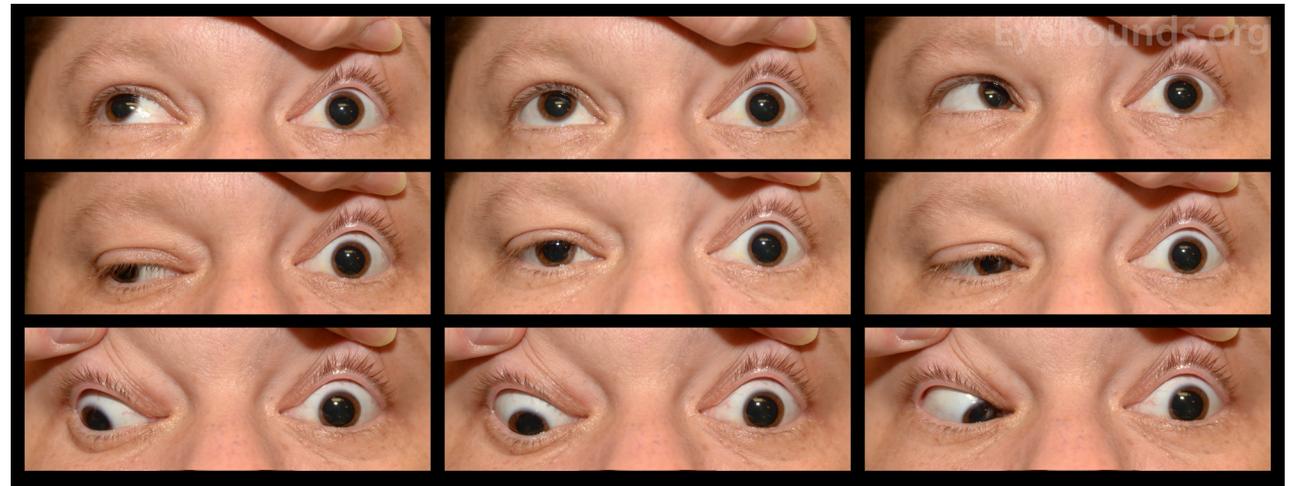
Left CN VI Palsy



Left CN III Palsy



Left Cavernous Sinus Syndrome



Normal Pituitary MRI



T1 post-contrast sagittal

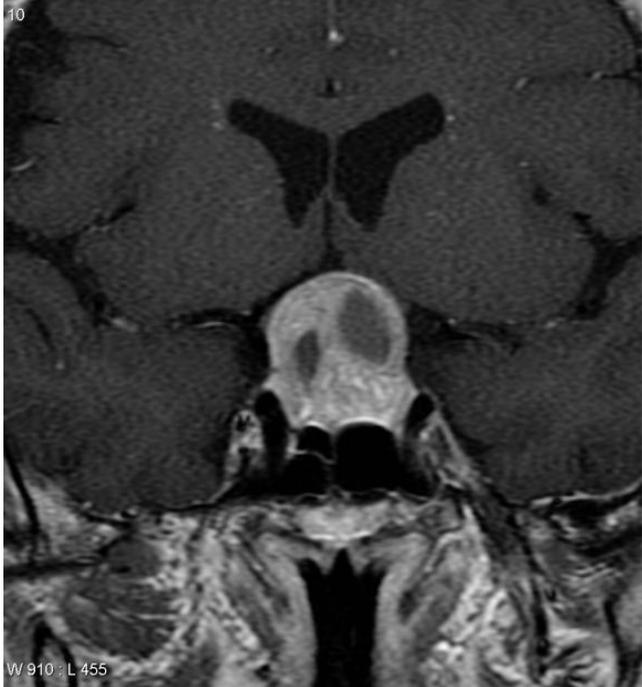
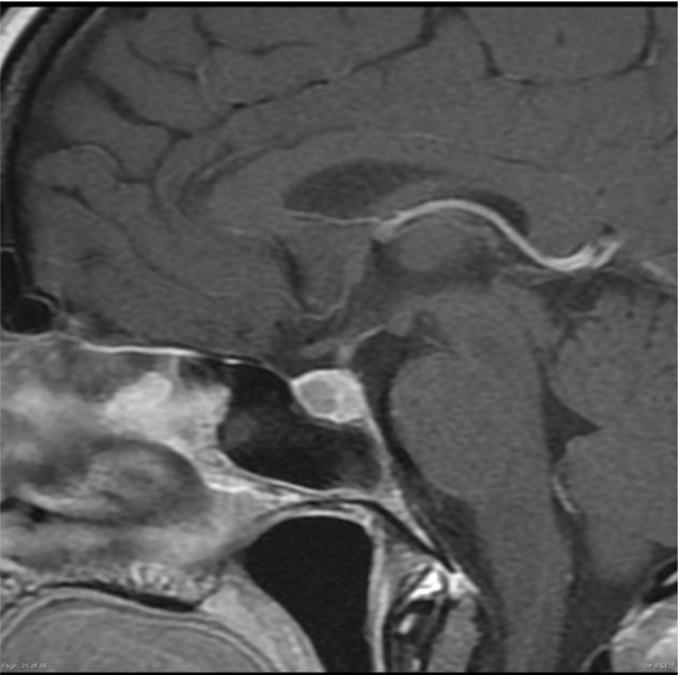
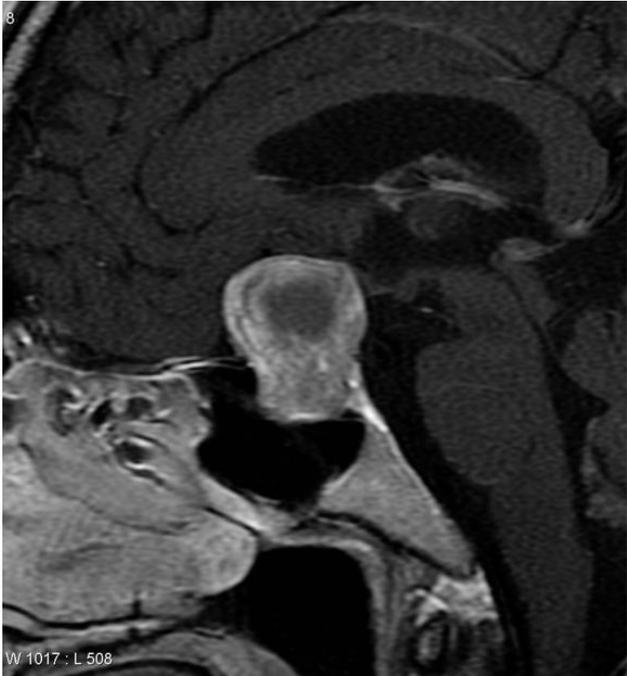


T1 post-contrast coronal

Microadenoma (<1cm)

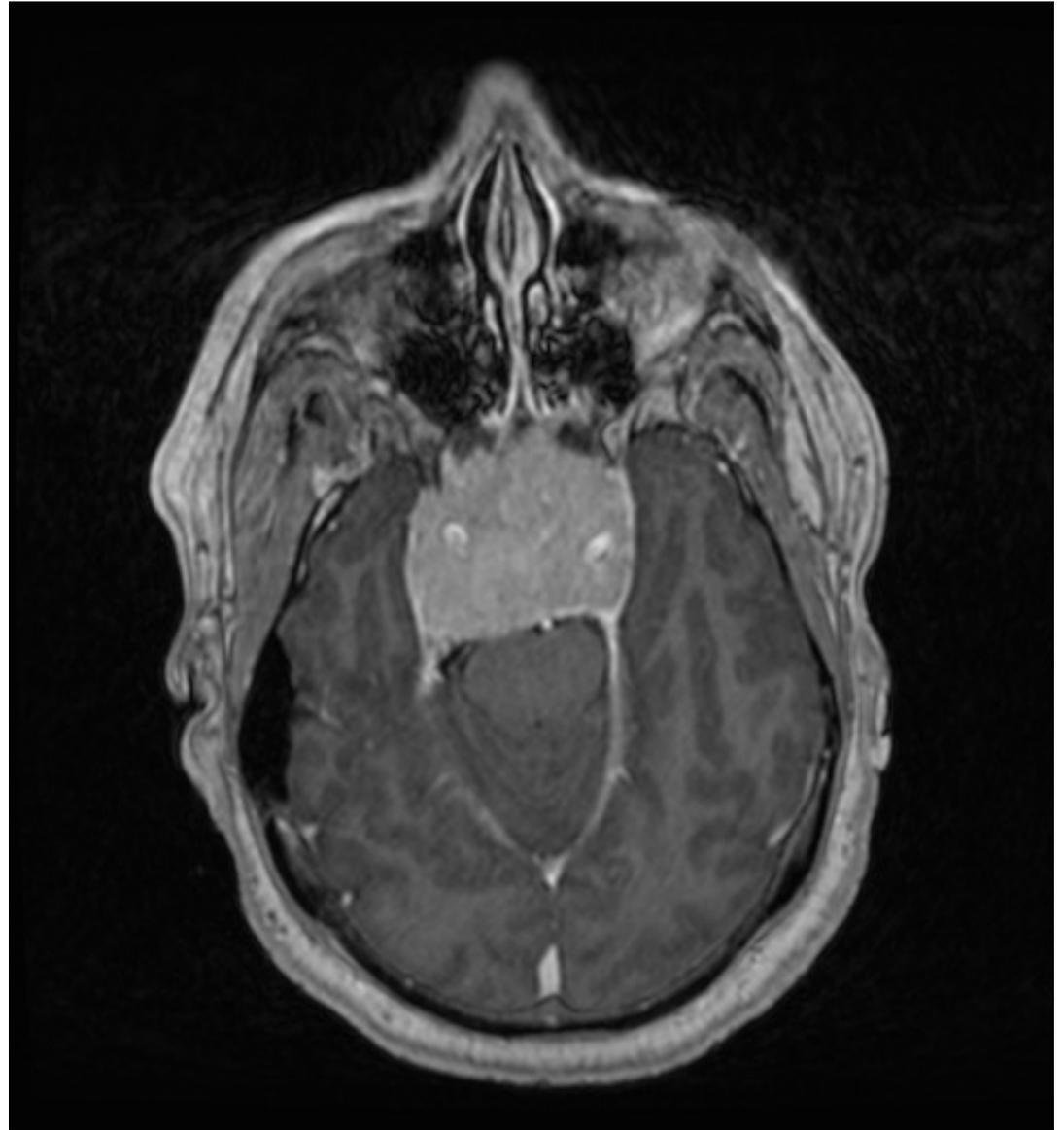
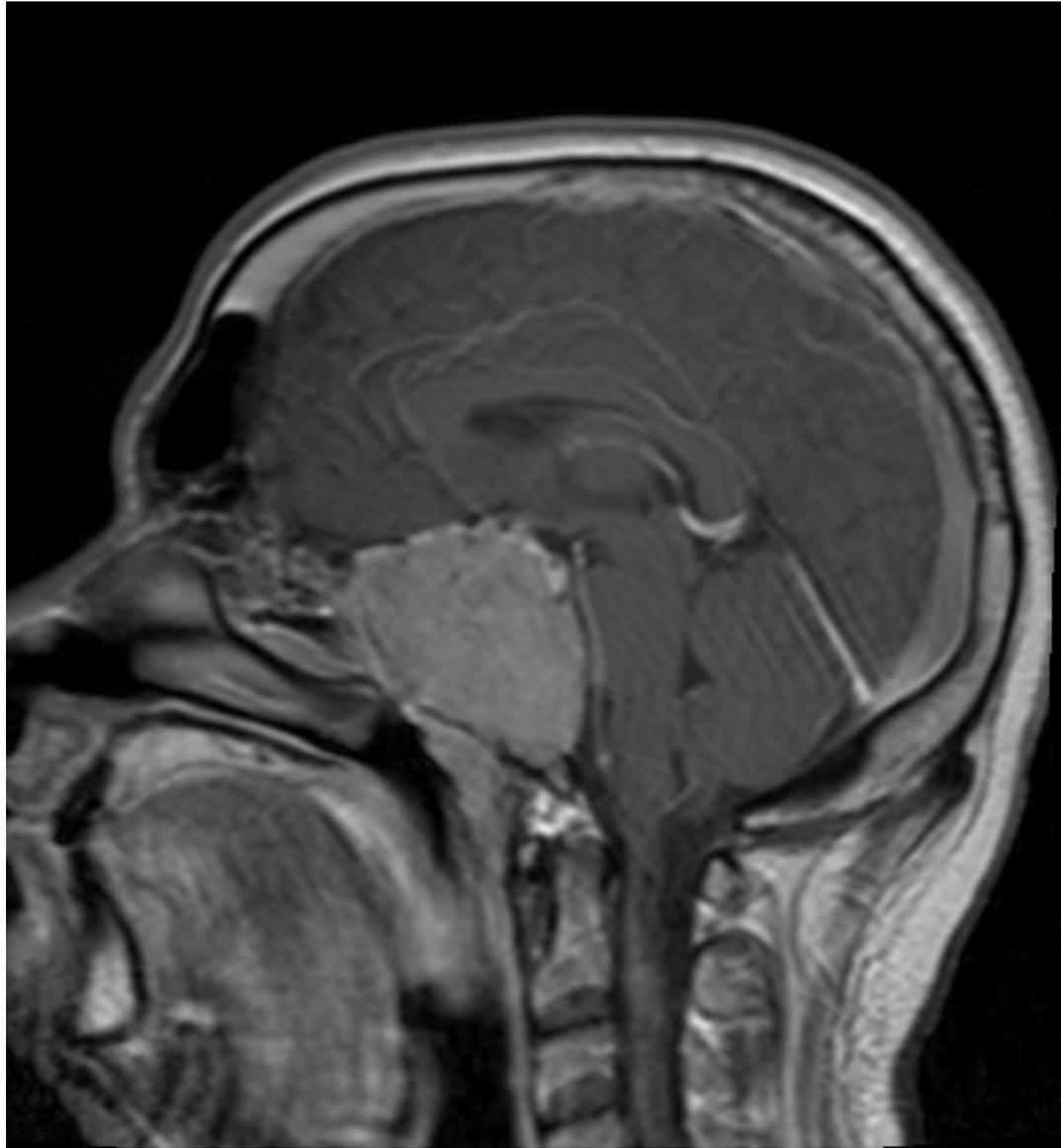


Macroadenoma (>1cm)



Functional Pituitary Adenomas

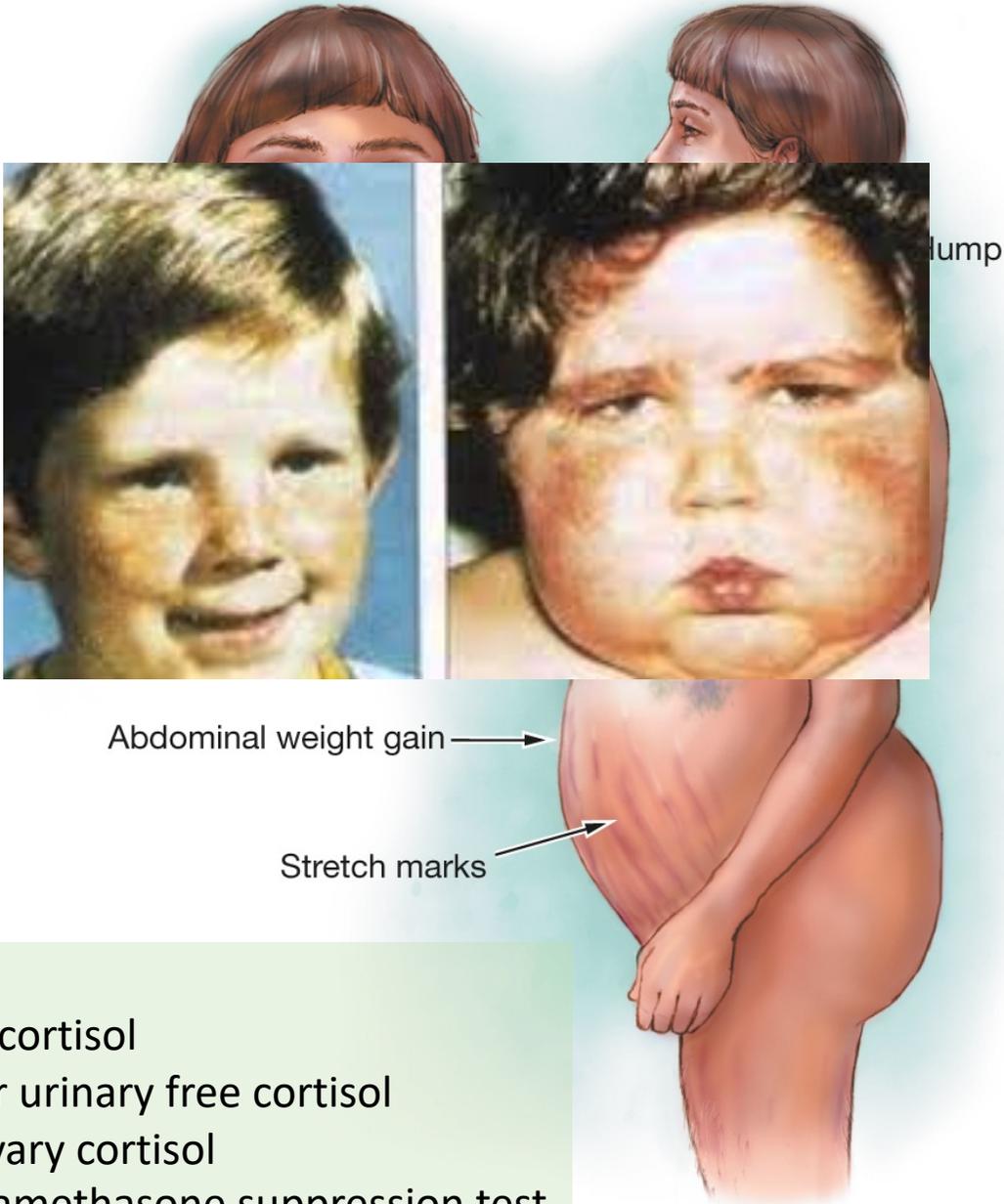
- Most commonly PRL (**Prolactinoma**)-> hyperprolactinemia
 - Females: amenorrhea, lactation
 - Males: erectile dysfunction
 - *PRL > 200 (< 200 = Stalk effect)*
 - Dilutions in clinically suspected but “low PRL” (Hook effect)
 - Treatment: Dopamine agonist (Cabergoline, bromocriptine)
- ACTH oversecretion: **Cushing’s disease**
 - Treatment: surgery
 - Other: radiation, medical.
- GH oversecretion: **Acromegaly**
 - “Surgical” disease
 - Medical management: octreotide, pegvisomant



PRL > 2000



Signs and symptoms of Cushing syndrome



- Dx:
- AM cortisol
 - 24hr urinary free cortisol
 - Salivary cortisol
 - Dexamethasone suppression test

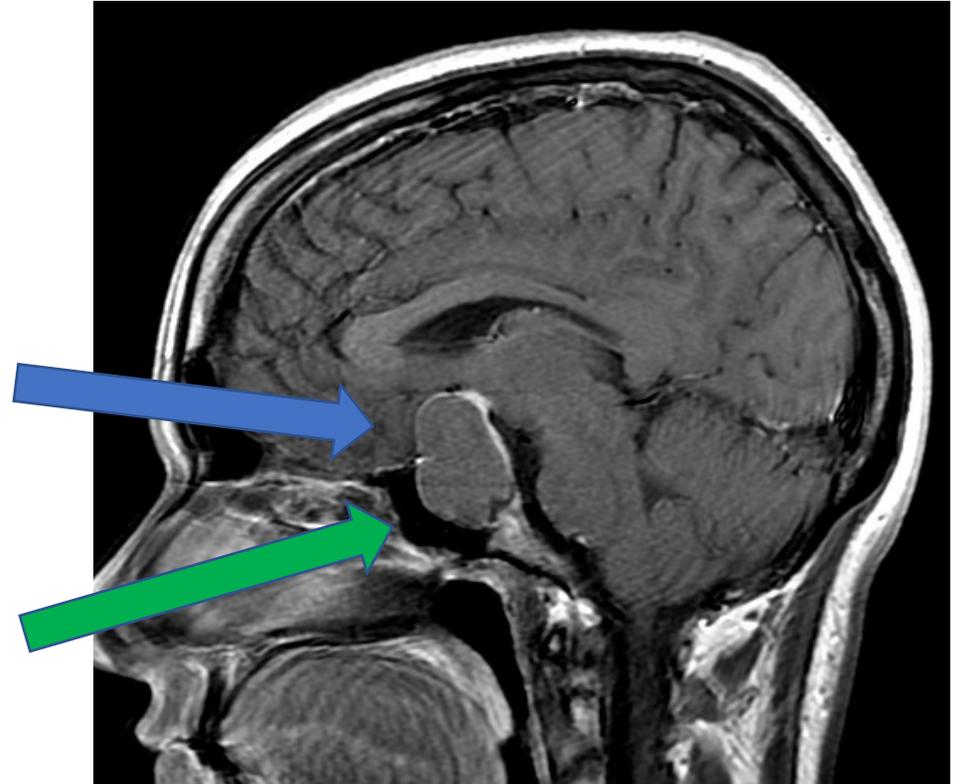
Acromegaly

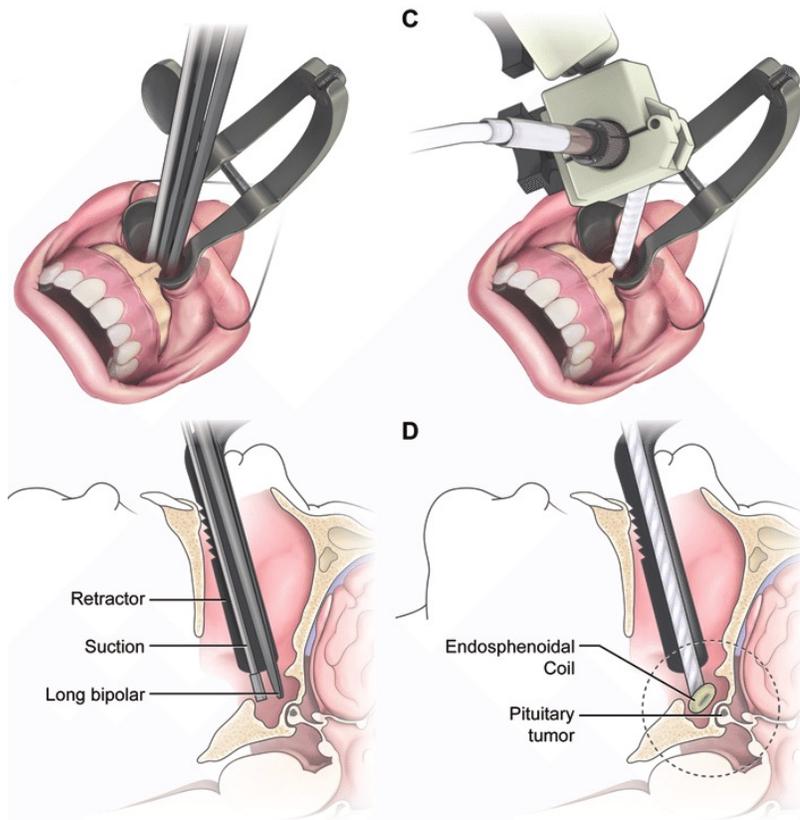


Dx: > IGF-1 and OGTT

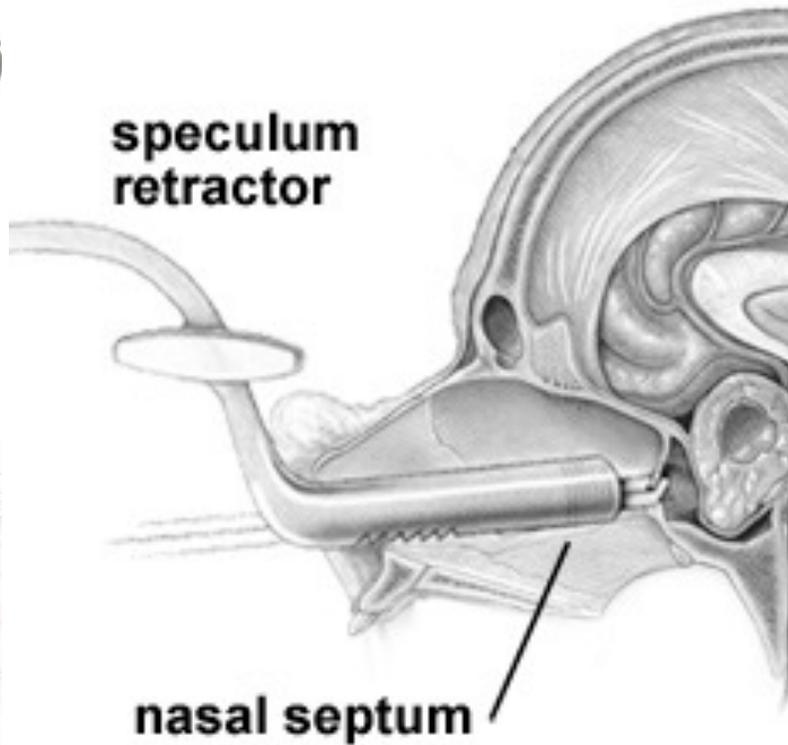
Pituitary Tumor Surgery

- Indications:
 - Vision loss / double vision
 - Prolactinoma refractory to Dopamine agonist therapy
 - Progressively enlarging on serial images.
 - Cushing's disease / Acromegaly
 - Endocrinopathies
- Approach: "Below or above"?
 - **Transphenoidal**
 - Transcranial

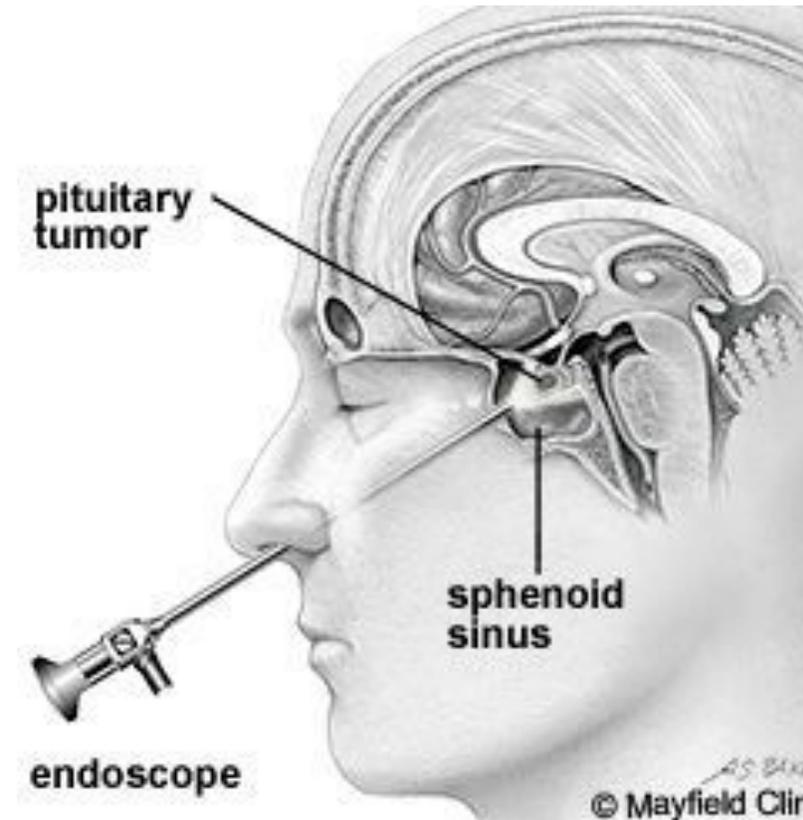




Sublabial Microscopic



Endonasal Microscopic



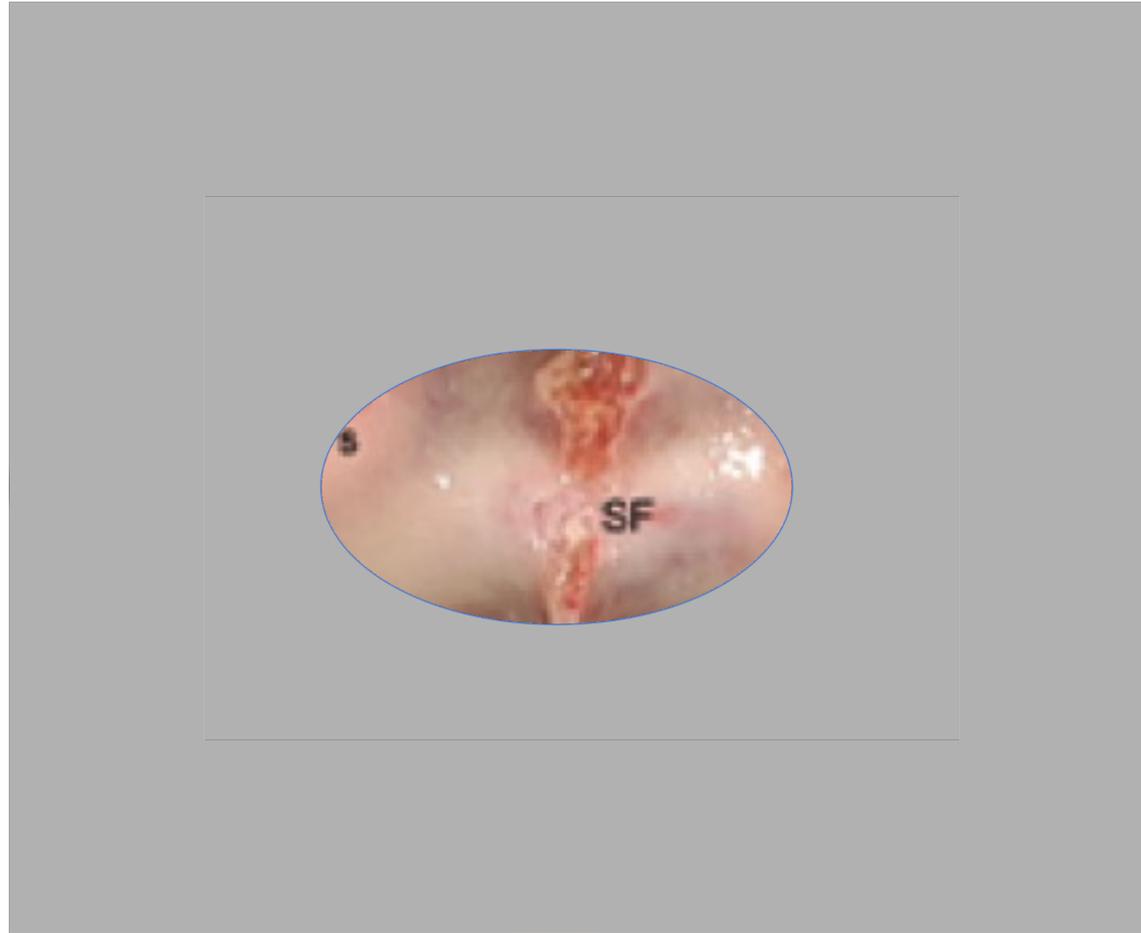
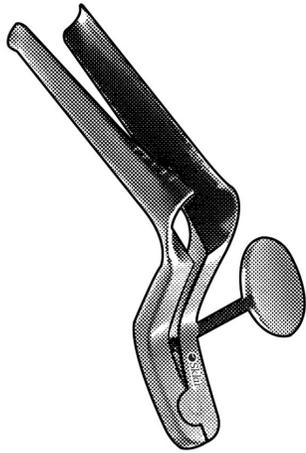
Endonasal Endoscopic

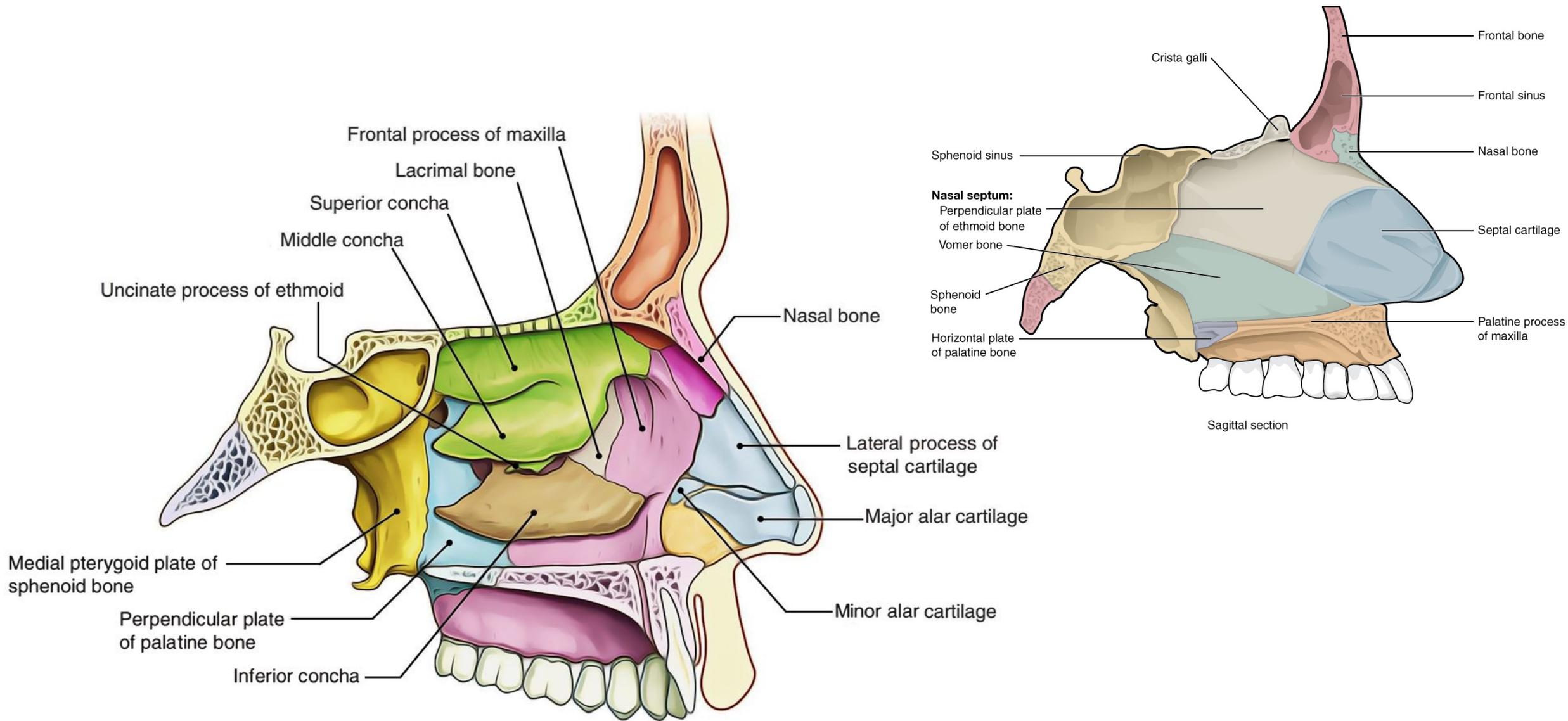
Transphenoidal Pituitary Surgery



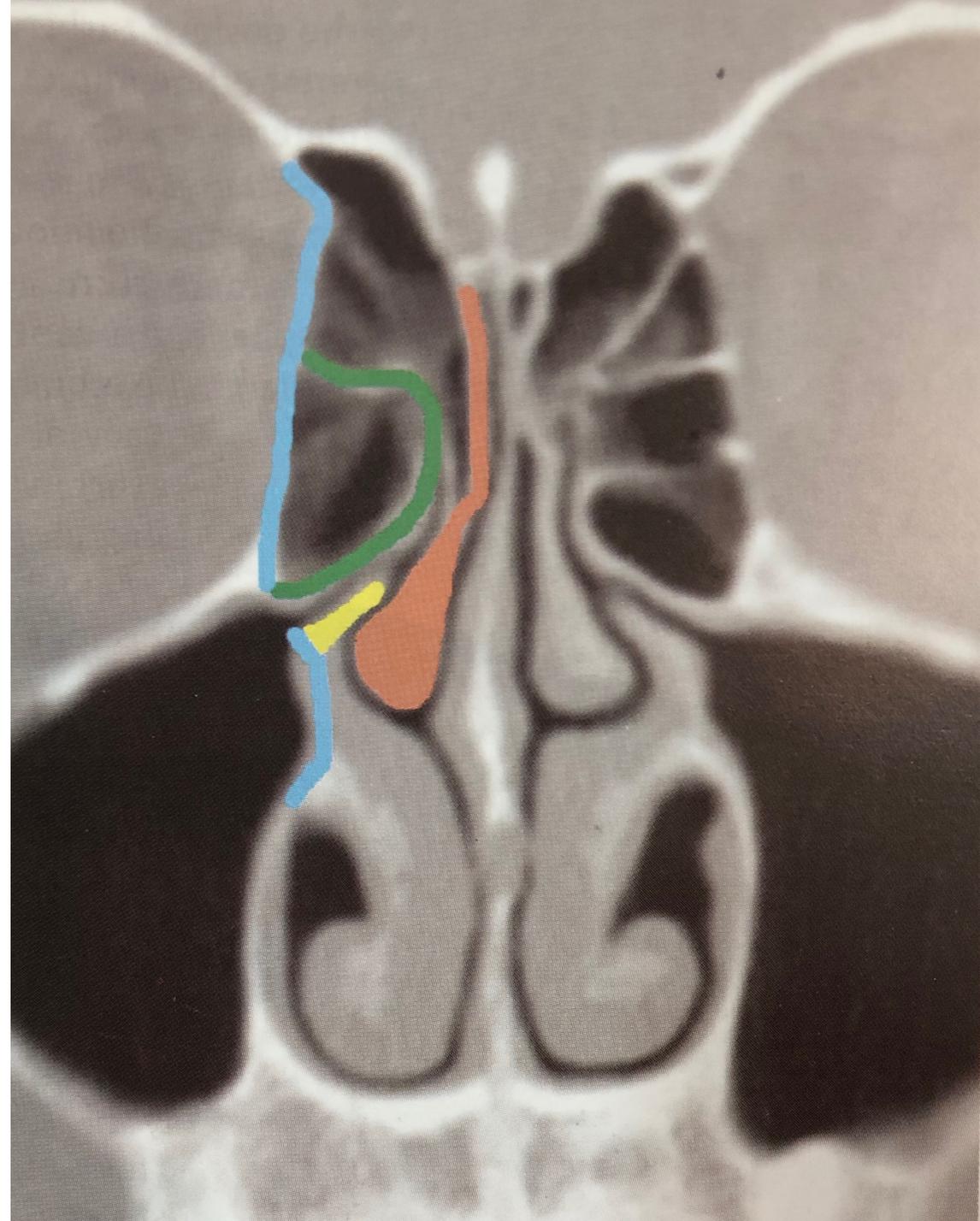
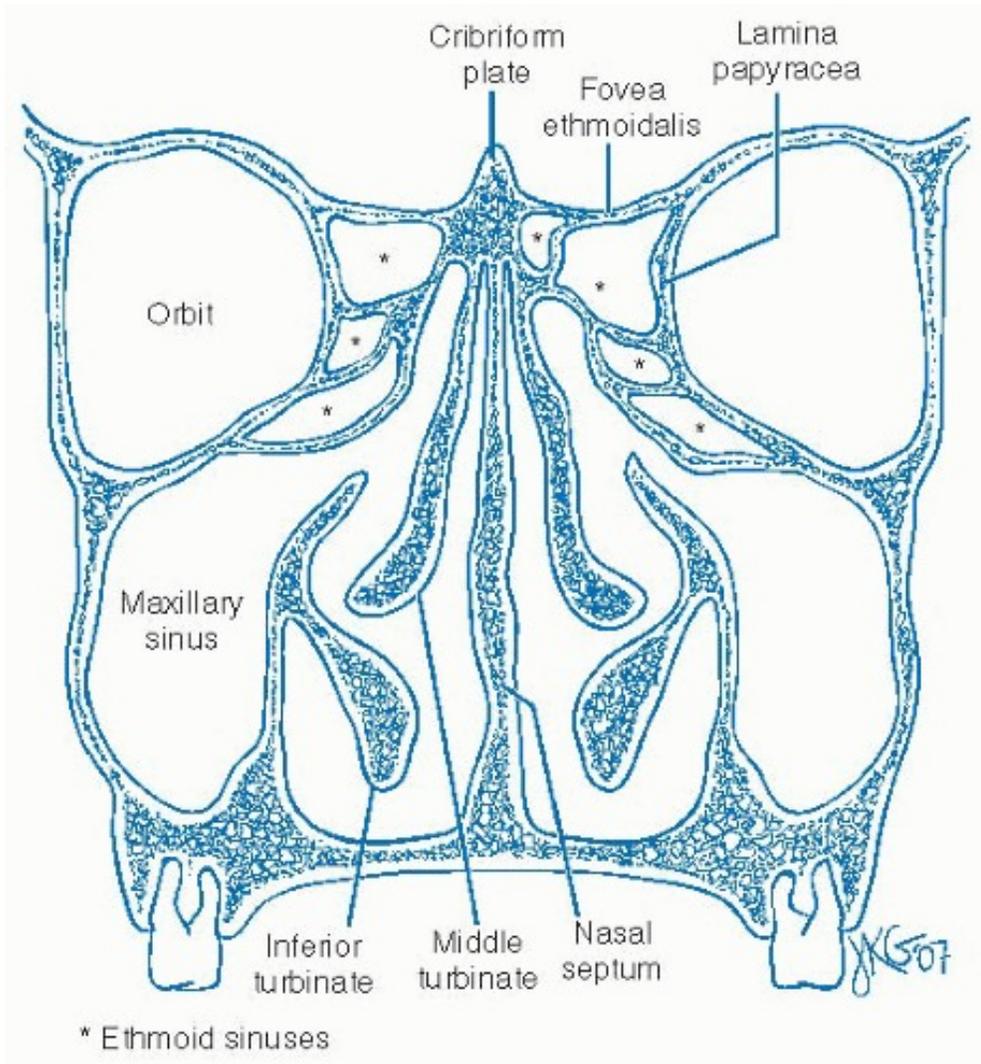
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Endoscopic vs Microscopic View

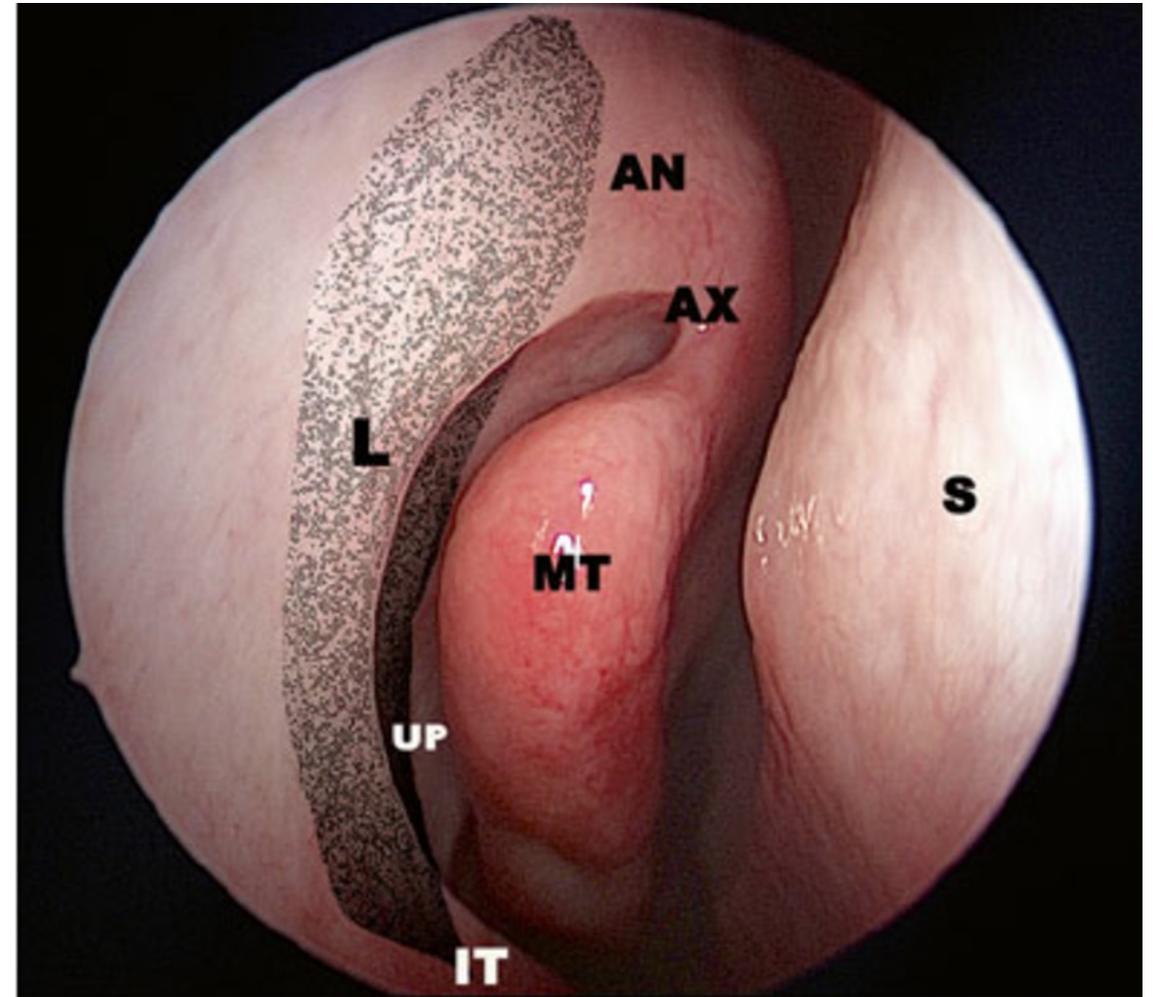
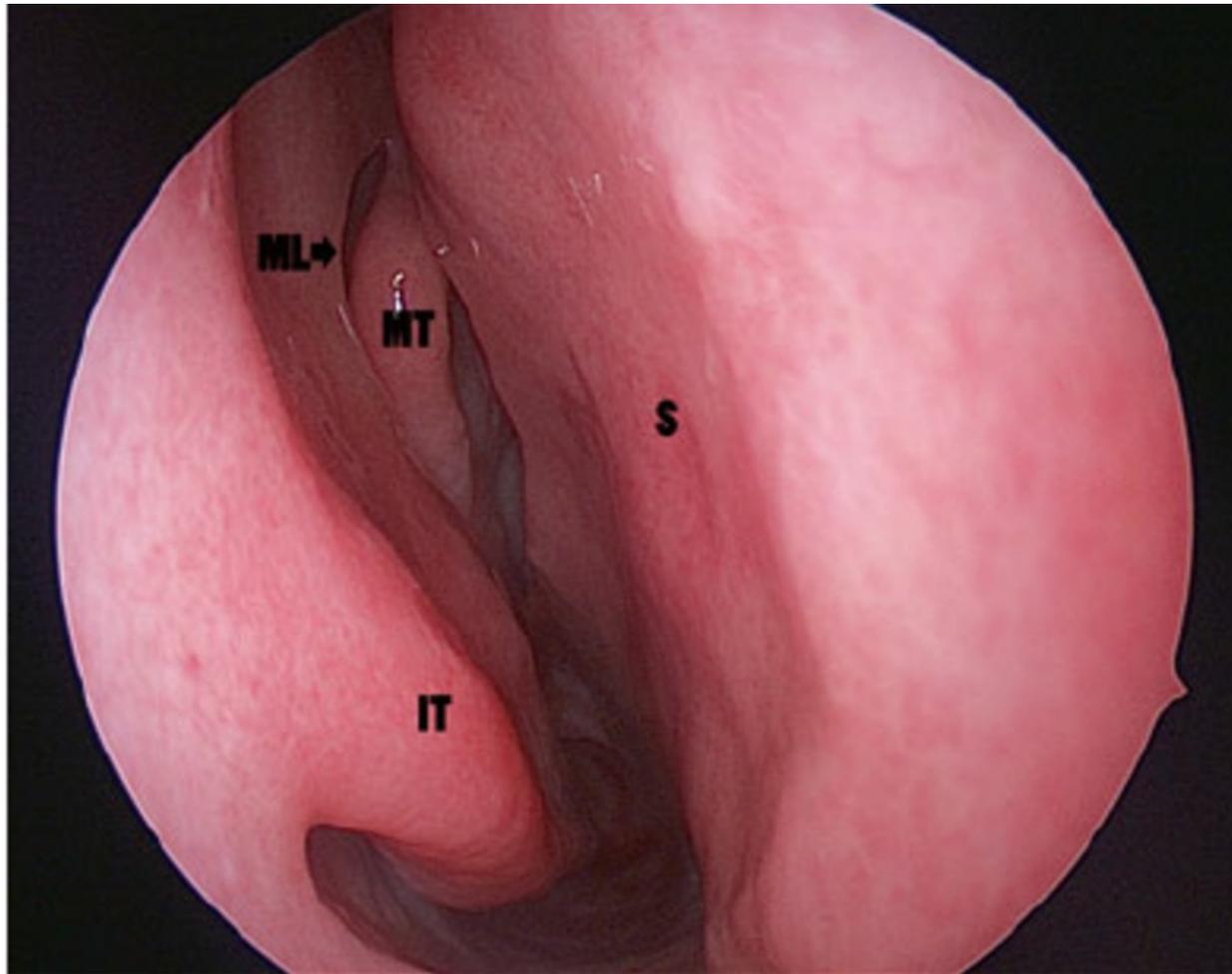




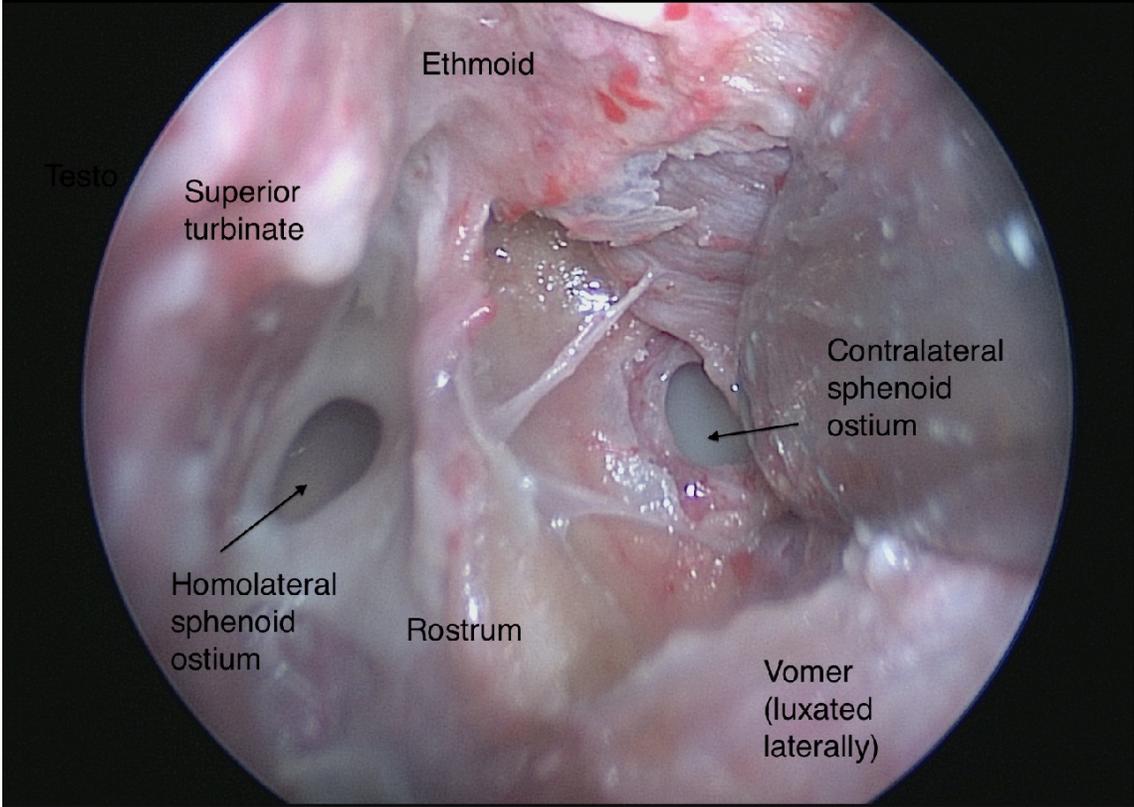
Sinonasal Cavity Anatomy – Lateral Nasal Wall and Nasal Septum



Ethmoid Bone & Turbinates

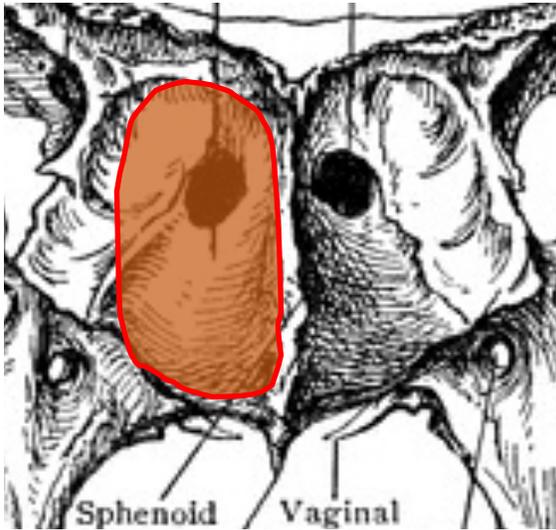


Sphenoidotomy Technique

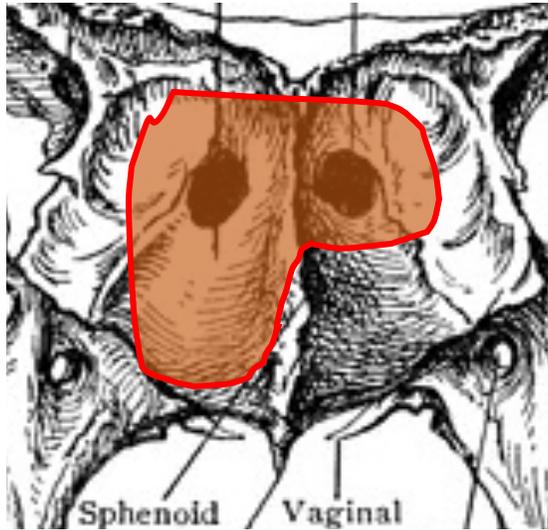


Approaches to the Sphenoid

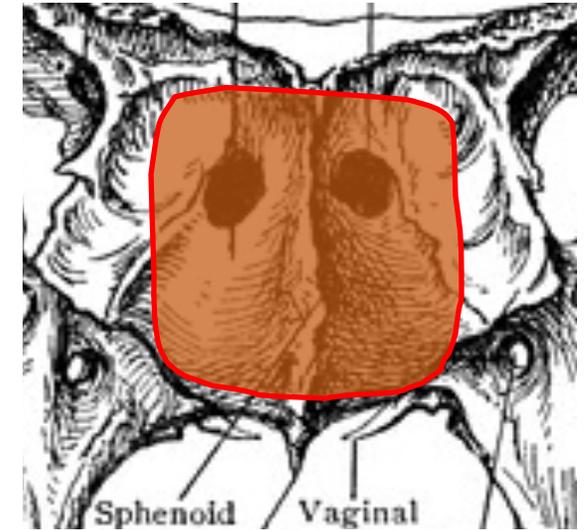
Unilateral



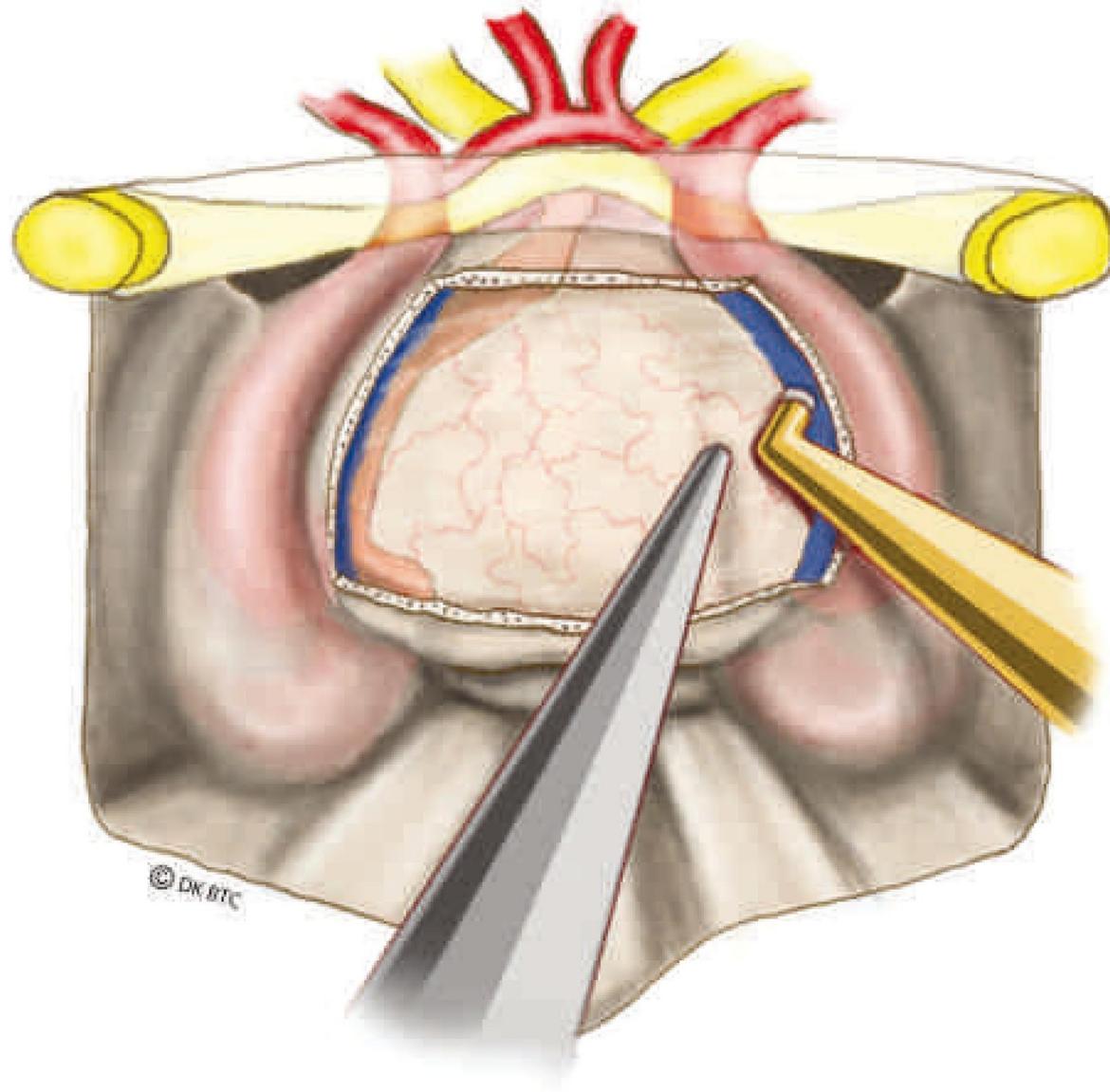
1.5 Approach



Bilateral Sphenoidotomies

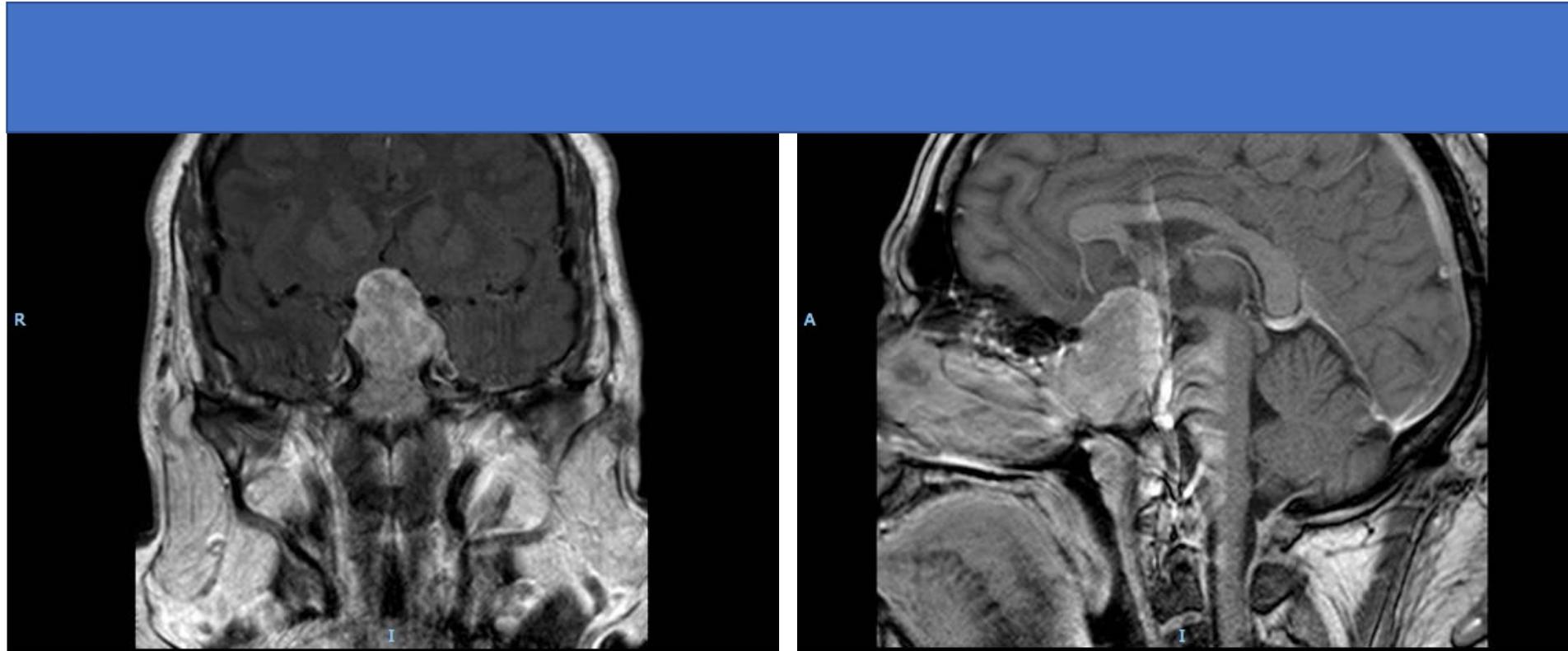


Thomas Jefferson Classification

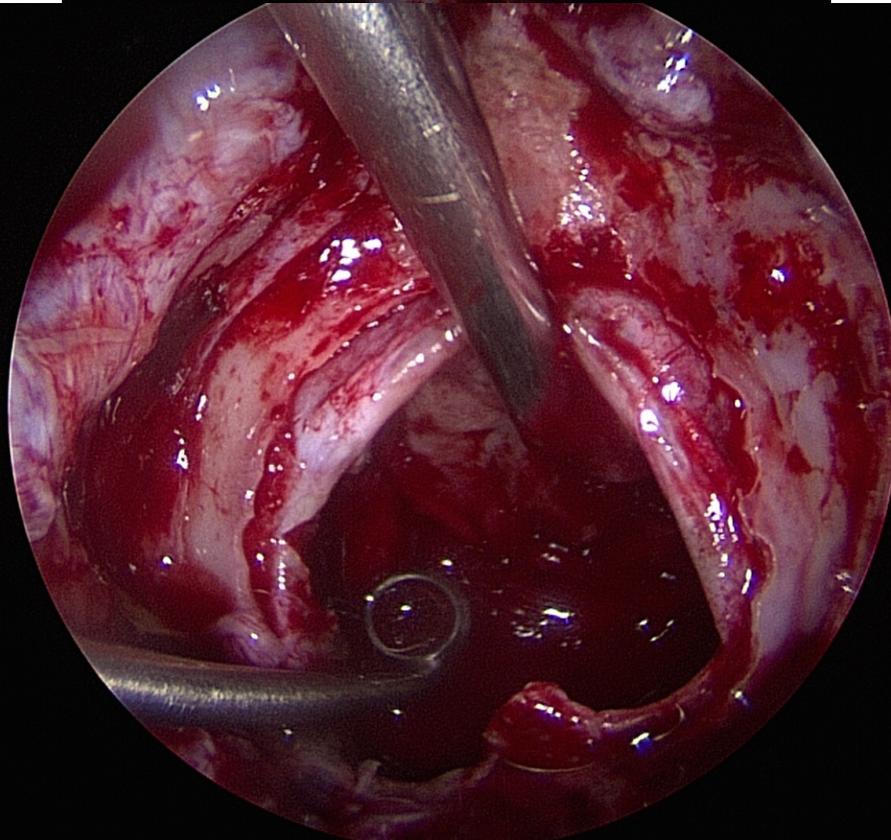
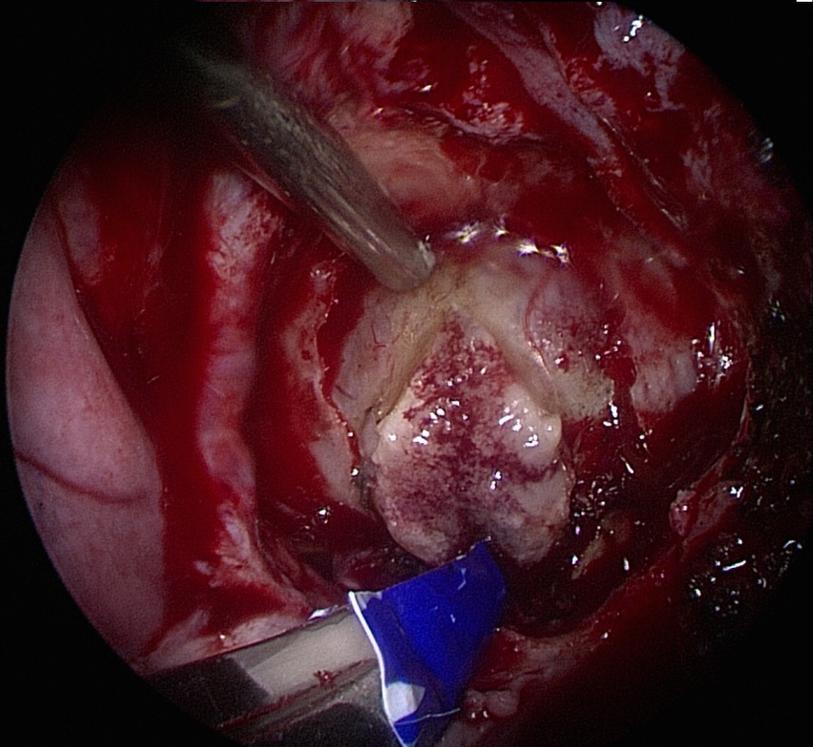
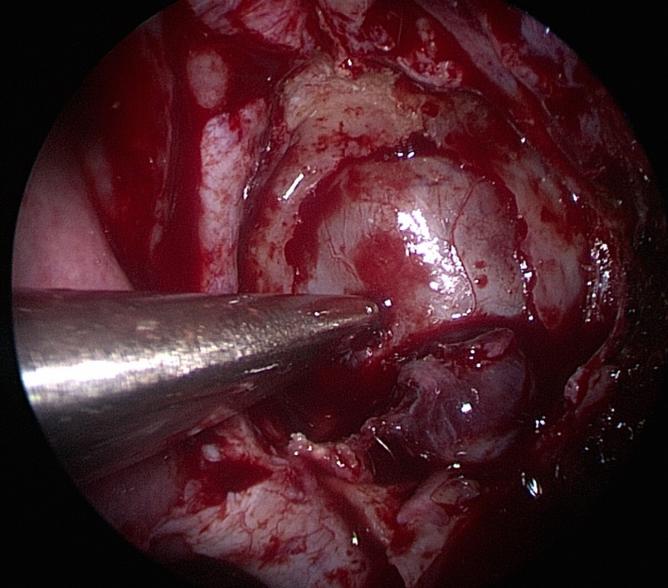
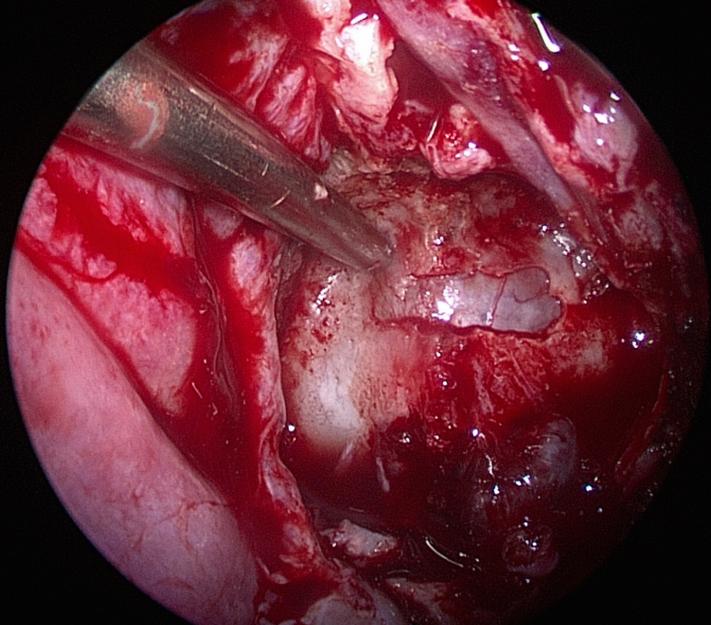


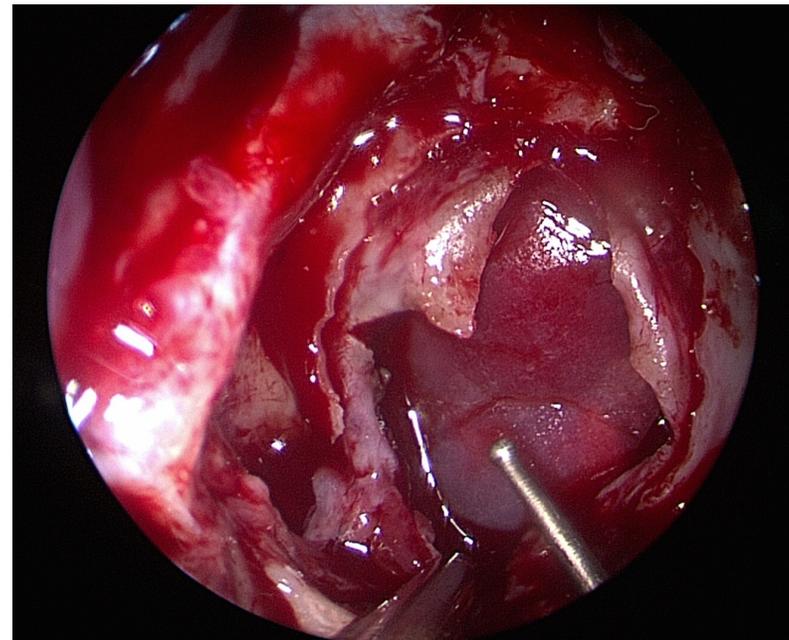
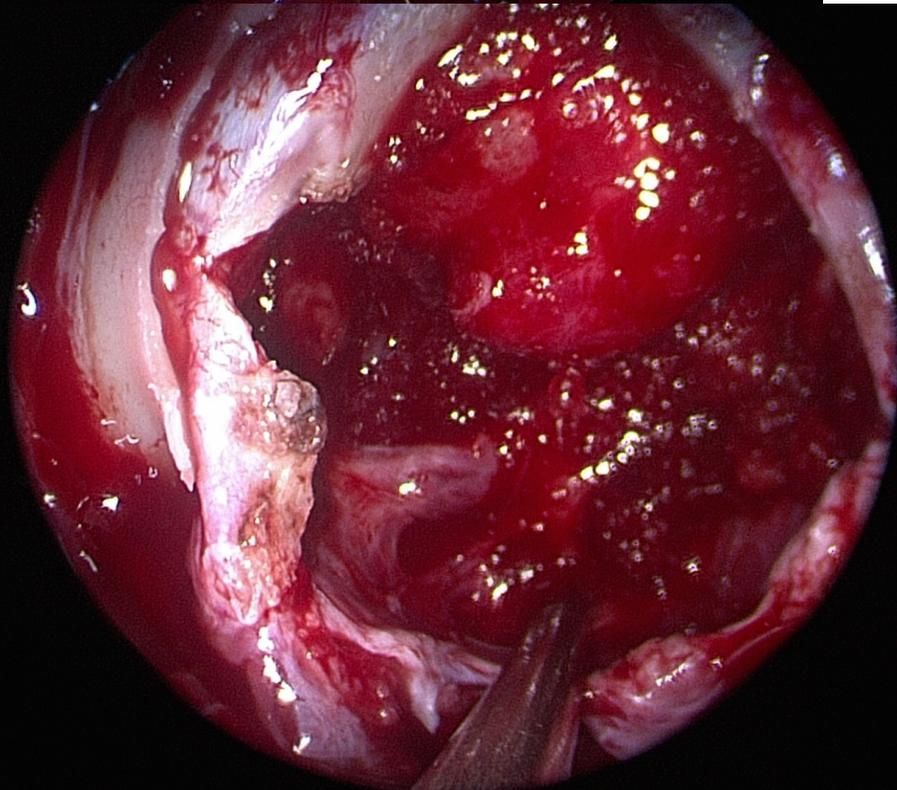
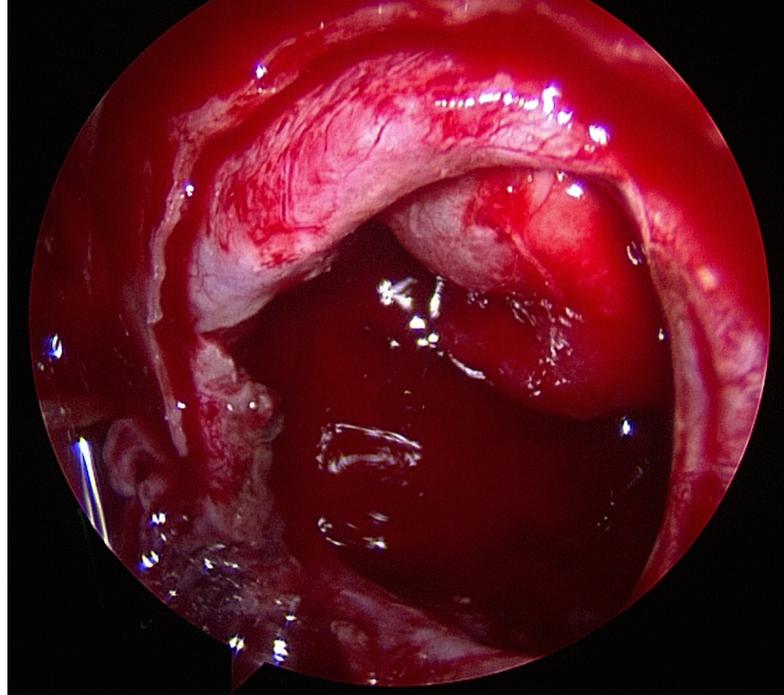
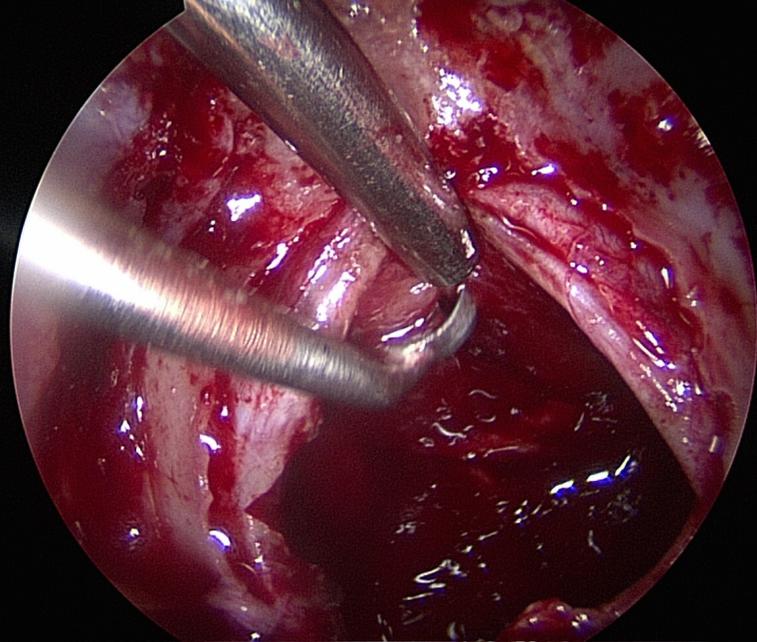
Sellar opening: CS to CS, up to tuberculum sellae.

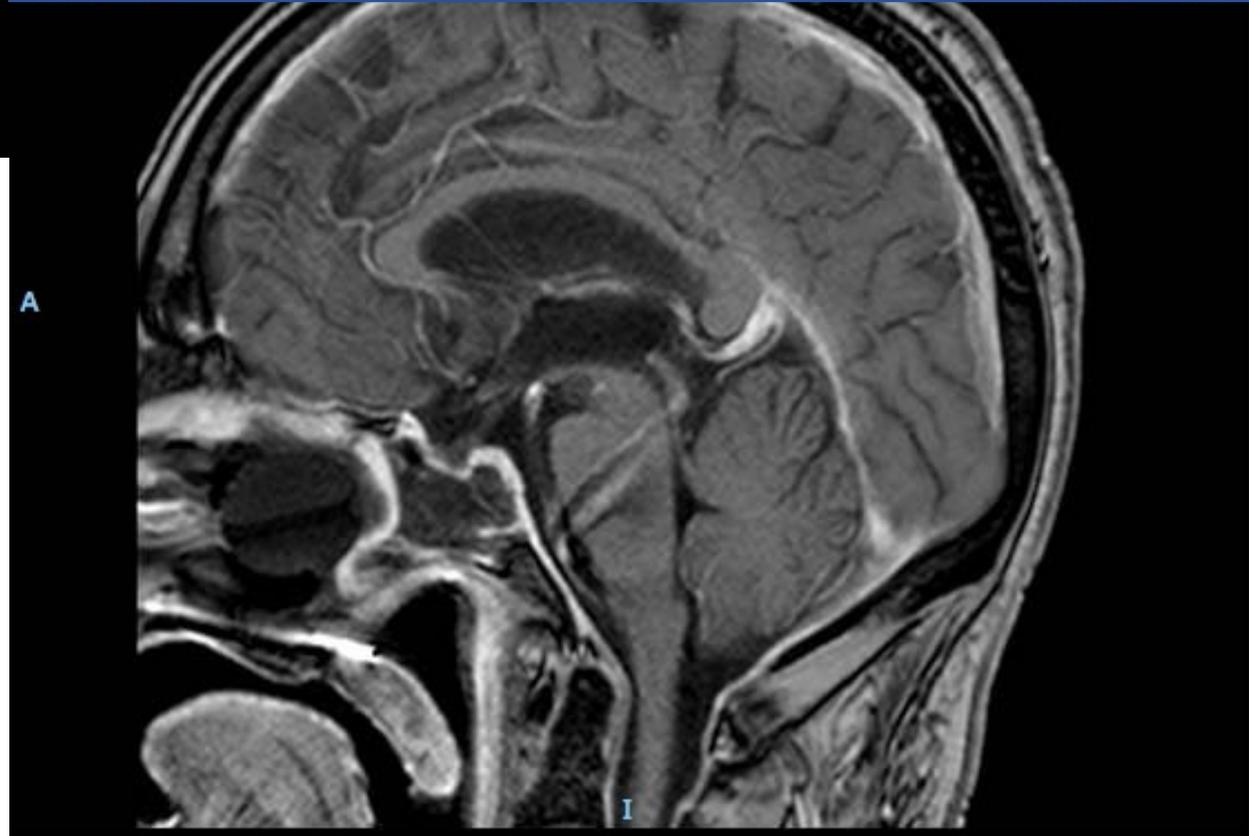
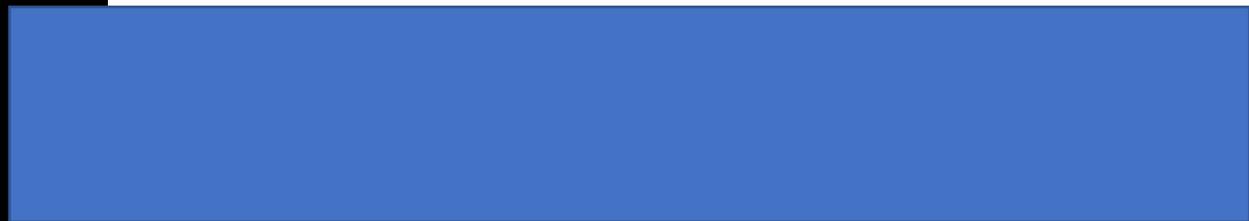
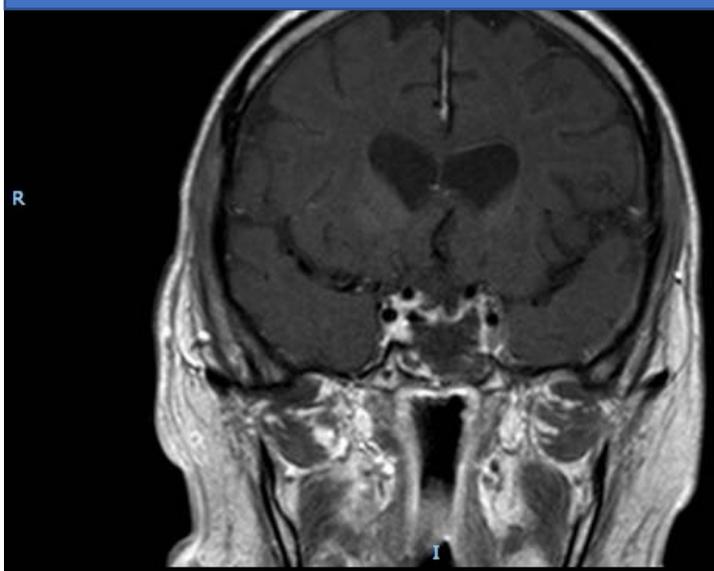
Pituitary Macroadenoma



44y/o M with Hx of vision loss and headaches. PRL and rest of hormones WNLs.

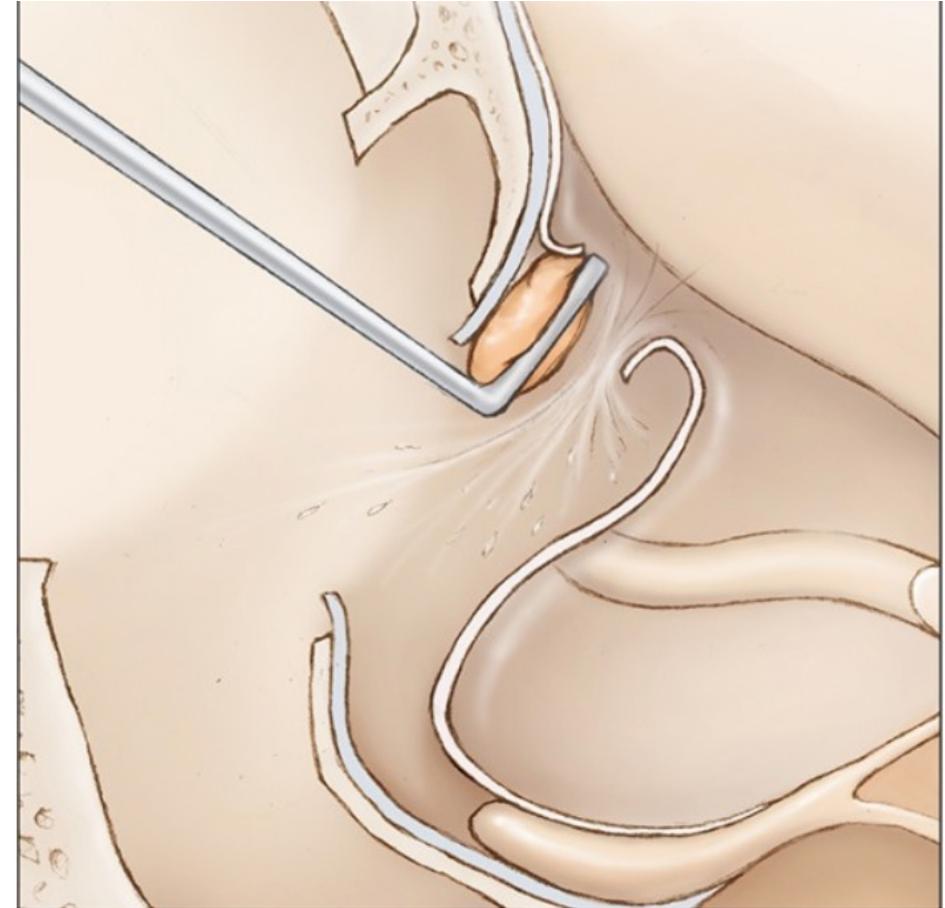






Sellar Closure

- No CSF leak, small dural defect: Surgical
- No CSF leak, patulous diaphragm: **Inlay dural repair + tissue sealant** (Tisseel, DuraSeal, Adherus)
- CSF leak:
 - Low flow and roof of sella: Inlay dural repair + tissue sealant
 - Fishmouth/tuberculum sella: Inlay graft with nasoseptal flap.

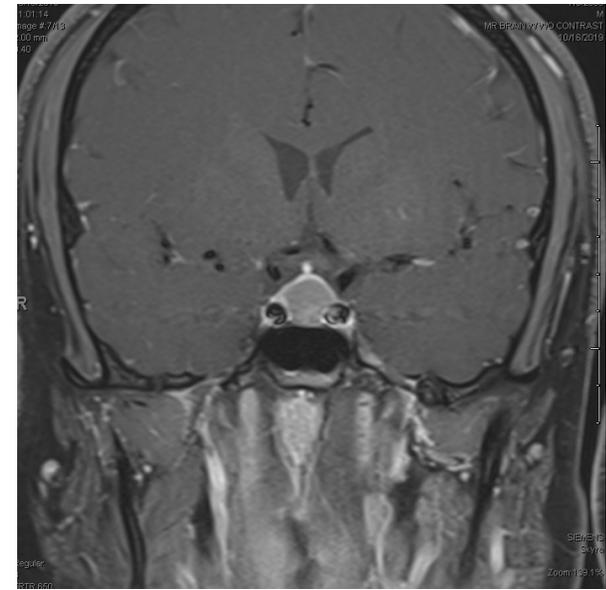
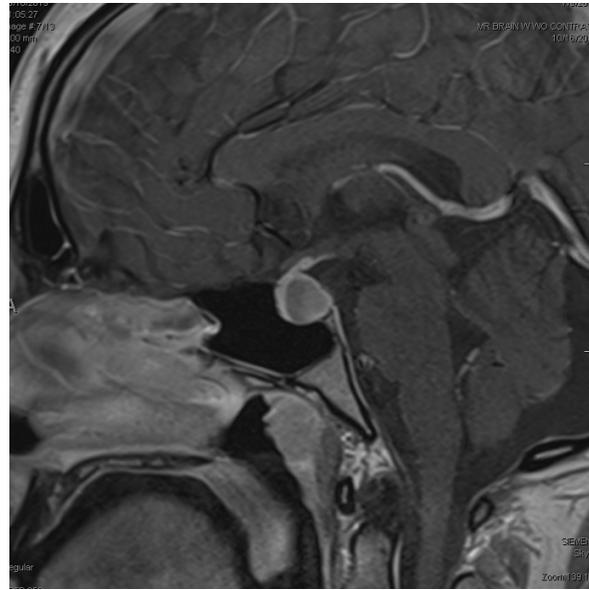
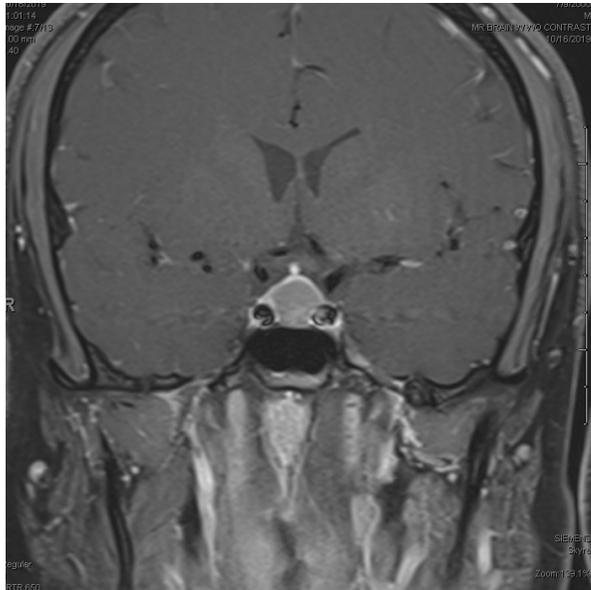


Post-Operative Care

- Neuro Step-Down
- HOB >45 degrees
- Ad lib PO H₂O intake at bedside.
- Strict I/Os
- CSF leak watch:
 - CSF rhinorrhea
 - Post-nasal drip
 - Head CT: parasellar pneumocephalus
- DI watch
 - UO: > 350 x 2hrs
 - Check Na and spec grav
 - Spec grav \leq 1.005
 - Rising Na level towards higher range of normal
 - DDAVP: nasal spray vs PO

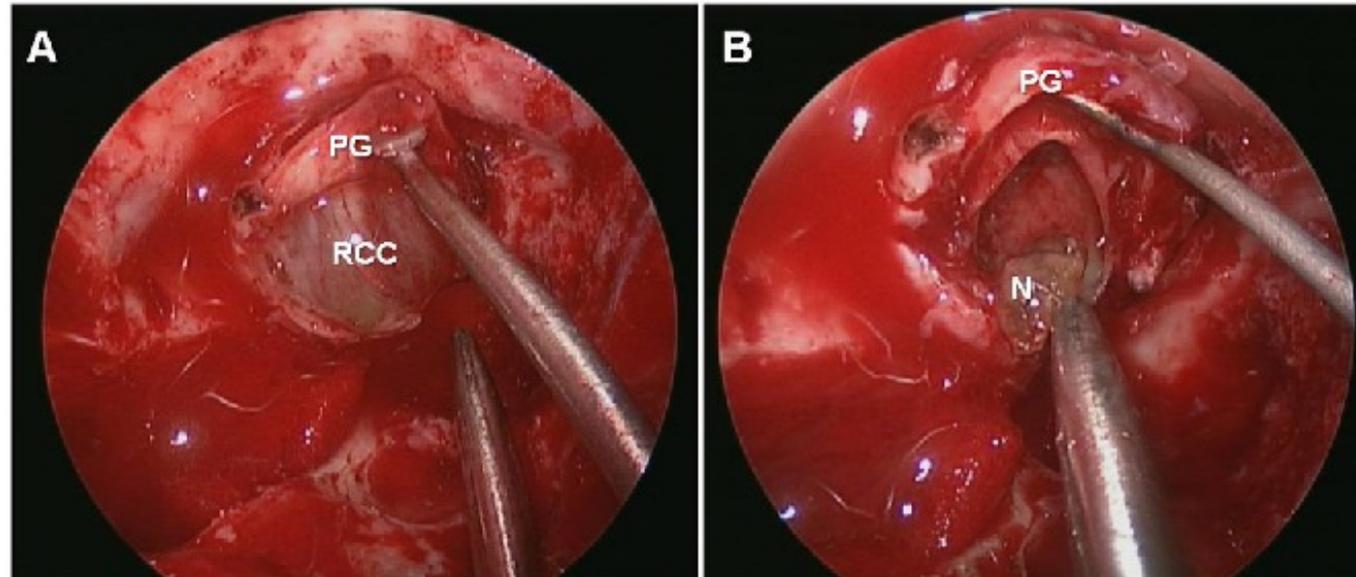
Rathke's Cleft Cyst

- Benign cystic remnants of the craniopharyngeal duct.
- Sellar & suprasellar region
- Majority of RCC's are discovered incidentally
- Sx's: H/As, endocrine dysfunction, visual loss; DI in 7-20% of patients



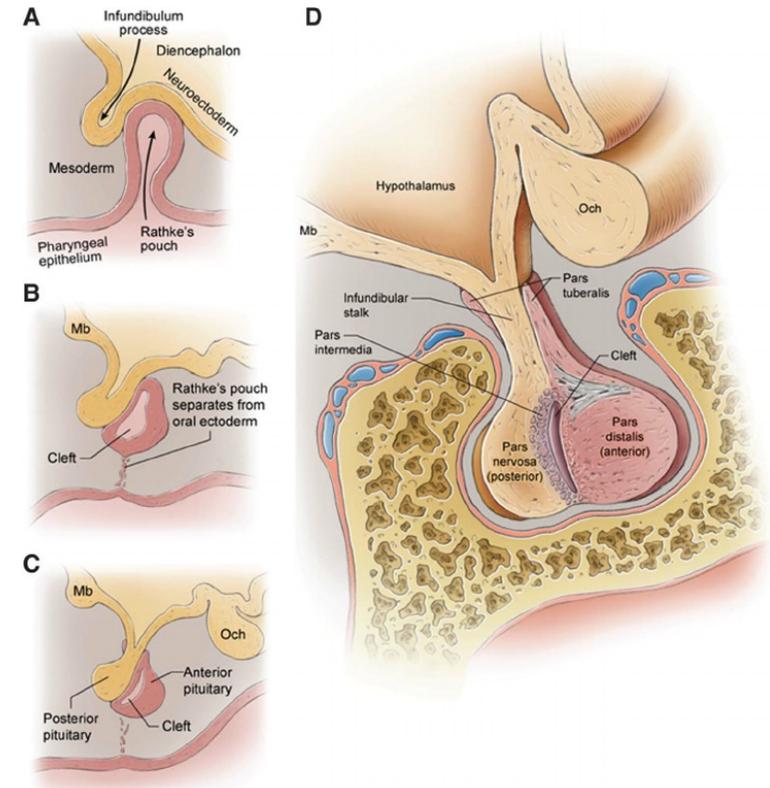
Rathke's Cleft Cyst Surgery

- Endoscopic transsphenoidal approach
- Wide fenestration and marsupialization of cyst
- Biopsy of wall prior to fenestration
- If CSF leak, repair as with pituitary adenoma.



Craniopharyngioma

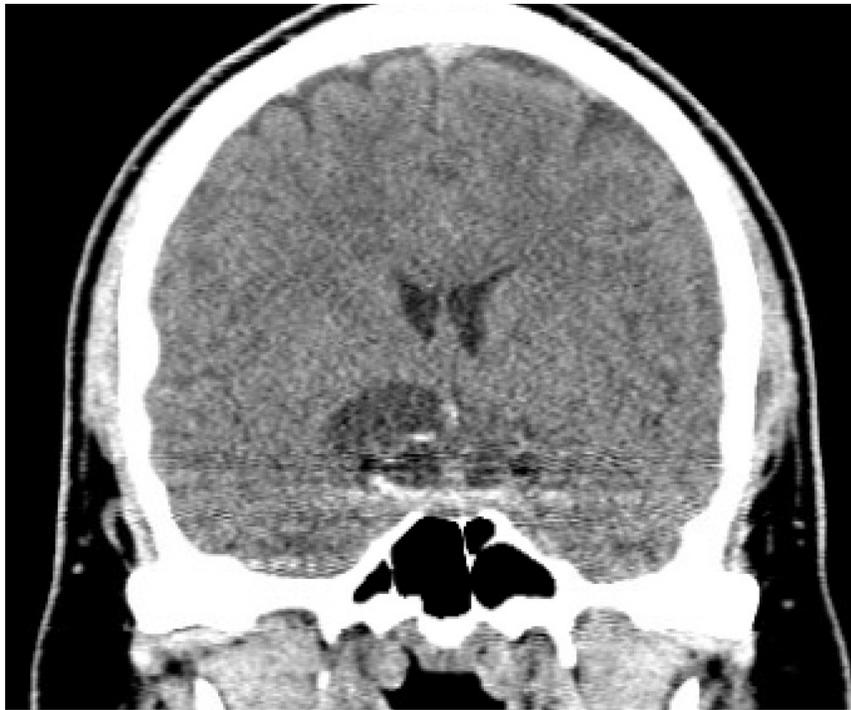
- Benign epithelial tumors that originate from the epithelial remnants of Rathke's pouch and the craniopharyngeal duct
 - WHO Grade I classification
- “Benign tumor in a malignant location”
- Local invasion of:
 - Hypothalamus
 - Optic chiasm
 - ICA, Pcom, hypophyseal arteries



“To be sure, one may occasionally succeed in stripping out a thin-walled cyst, and examples of this have been reported, but when the tumor is partly solidified and calcareous, sad experience warns the surgeon to leave it pretty much alone” –Harvey Cushing, 1932.

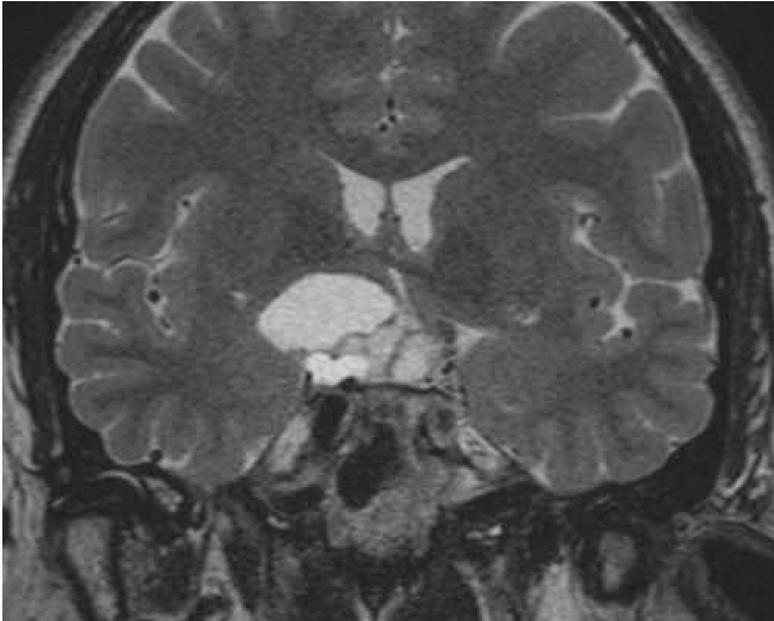
Radiographic Findings

- CT

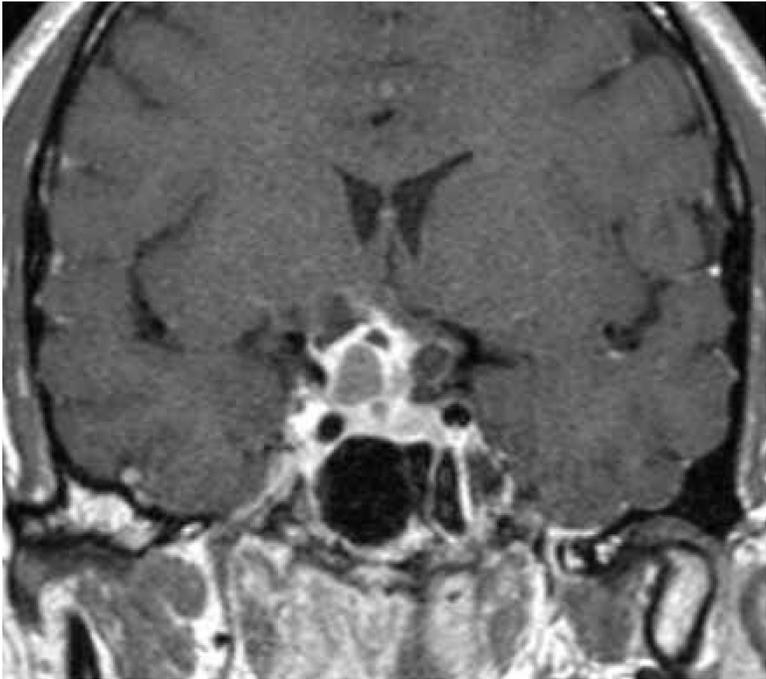
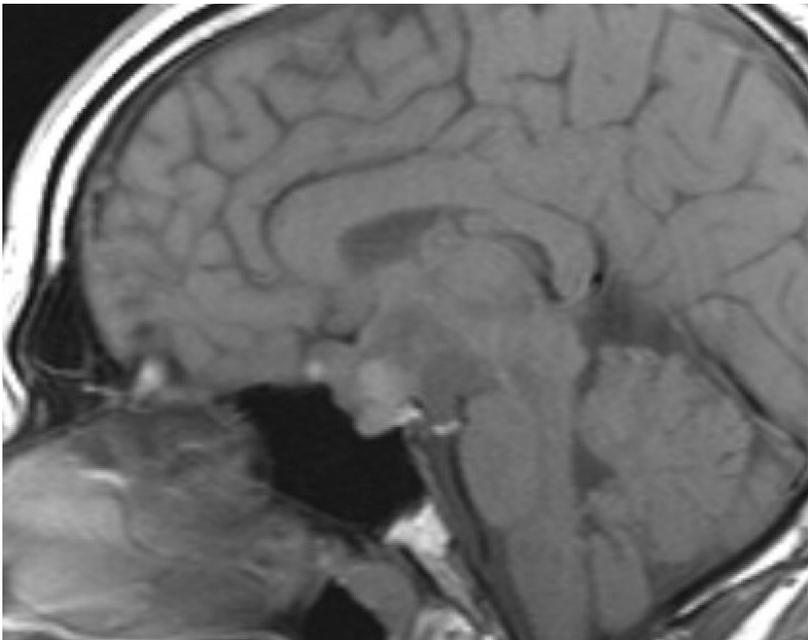


• MRI

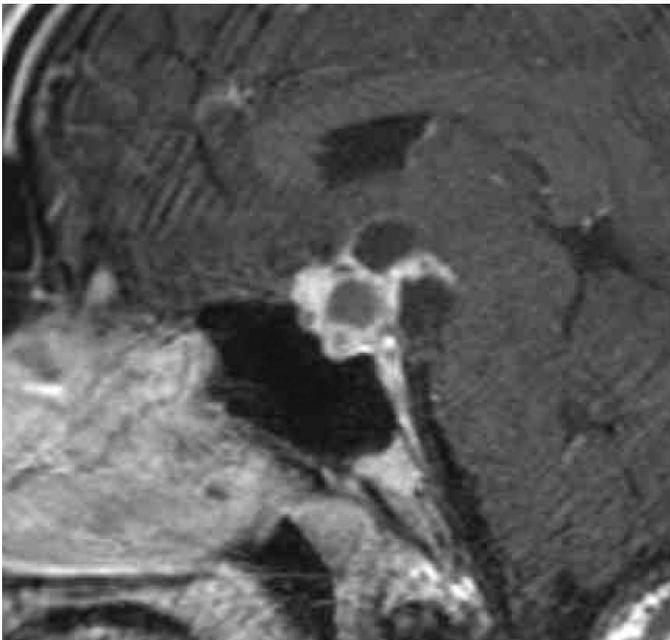
T2



T1 - C



T1 + C



T1 + C

Craniopharyngioma Treatment

- **Surgical resection**
 - **Transcranial approach**
 - **Transphenoidal: expanded endoscopic approach (EEA)**
- Stereotactic radiotherapy
- Intra-tumoral radioactive isotopes
- Intracystic catheter placement for drainage
- Hormone supplementation

Expanded Endoscopic Approach

Advantages

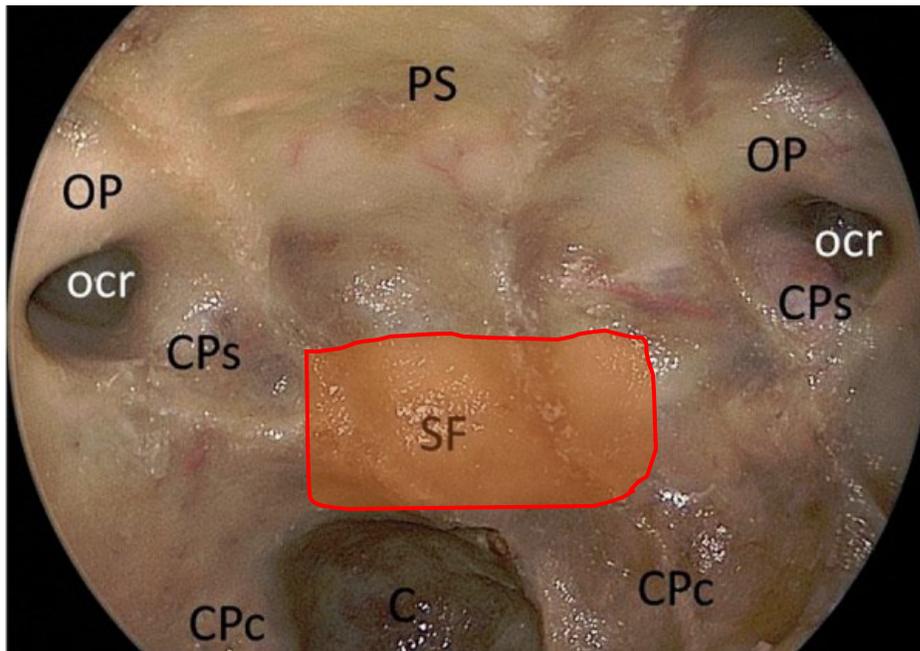
- "Minimally invasive".
- Midline approach for a midline lesion
- No brain retraction.
- Better preserve chiasmal perforators.
- Better visual preservation

Limitations

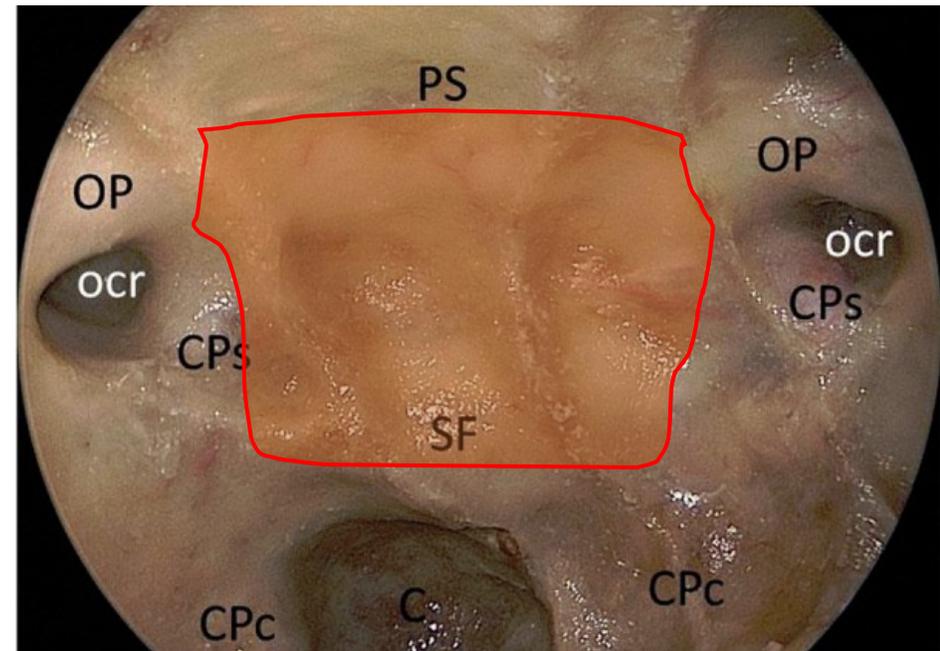
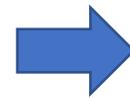
- Steep learning curve
 - Extensive experience in endoscopic transsphenoidal surgery
 - Fellowship training
- Lack of space and/or 3D perception
- Difficult to reach lateral components
- CSF leak

Expanded Endoscopic Transphenoidal Approach

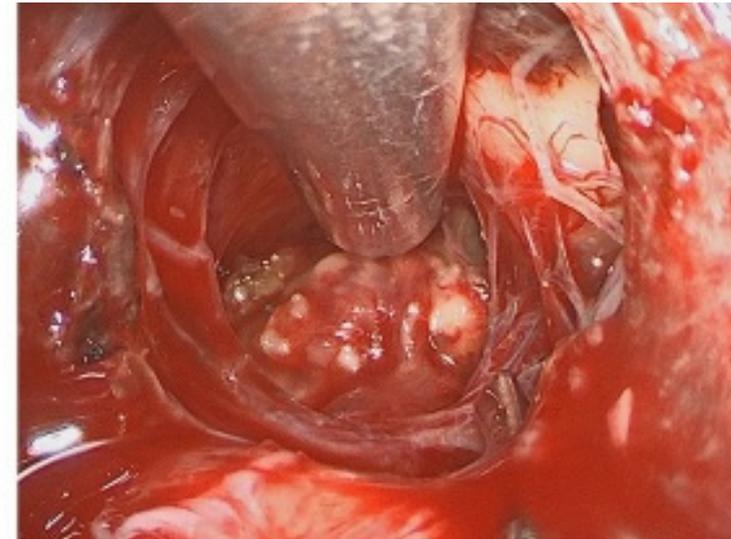
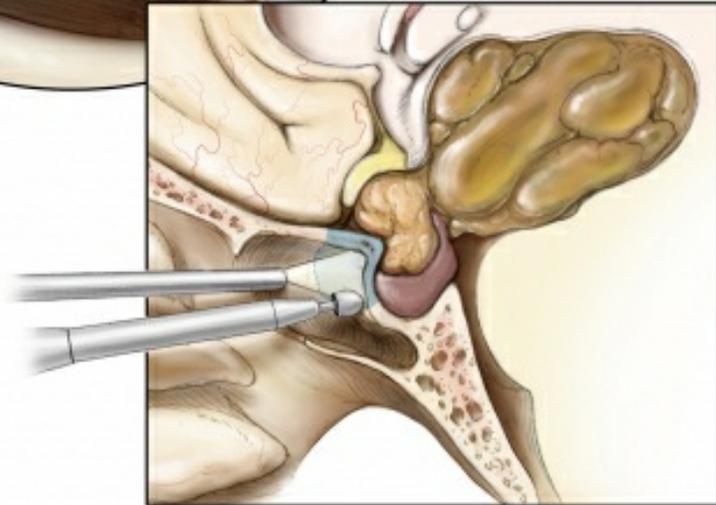
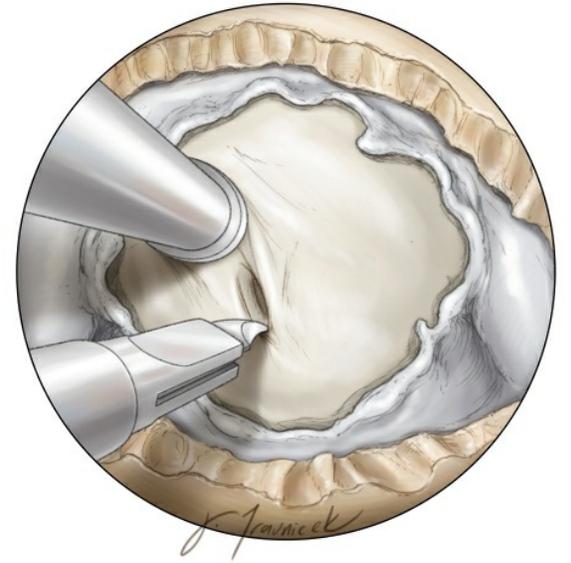
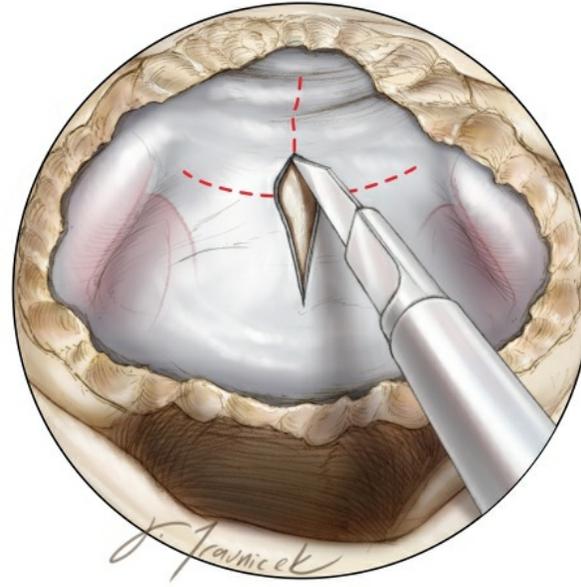
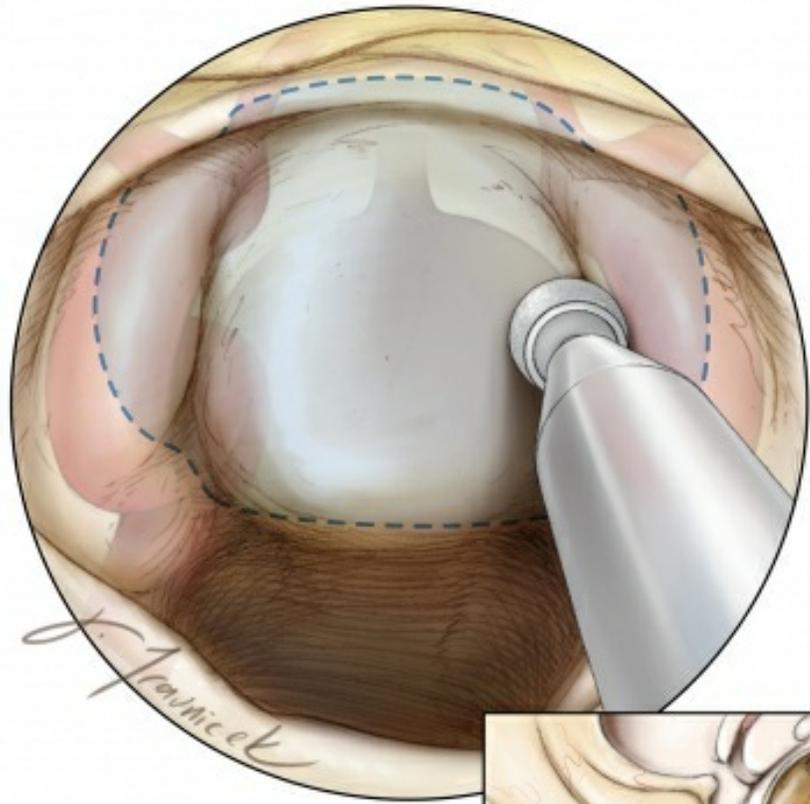
- Sellar approach + skull base removal (craniotomy)
- Craniopharyngioma: -> tuberculum sellae

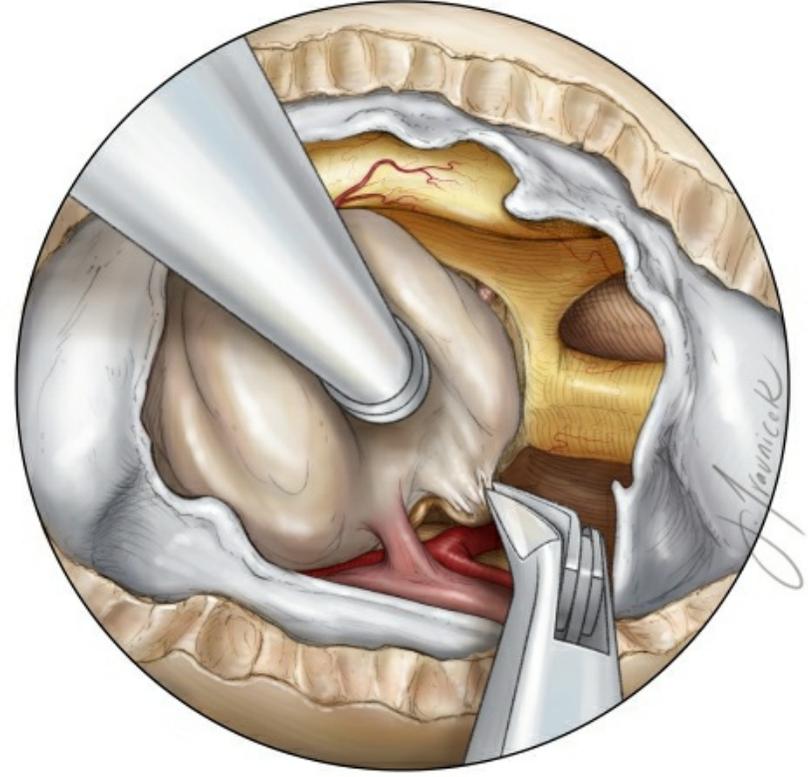
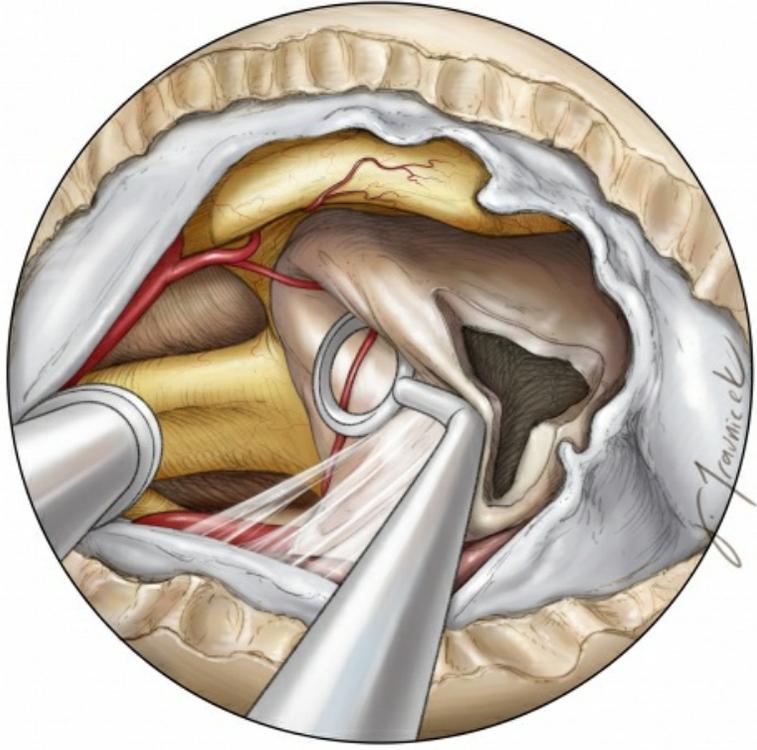


Sellar Approach



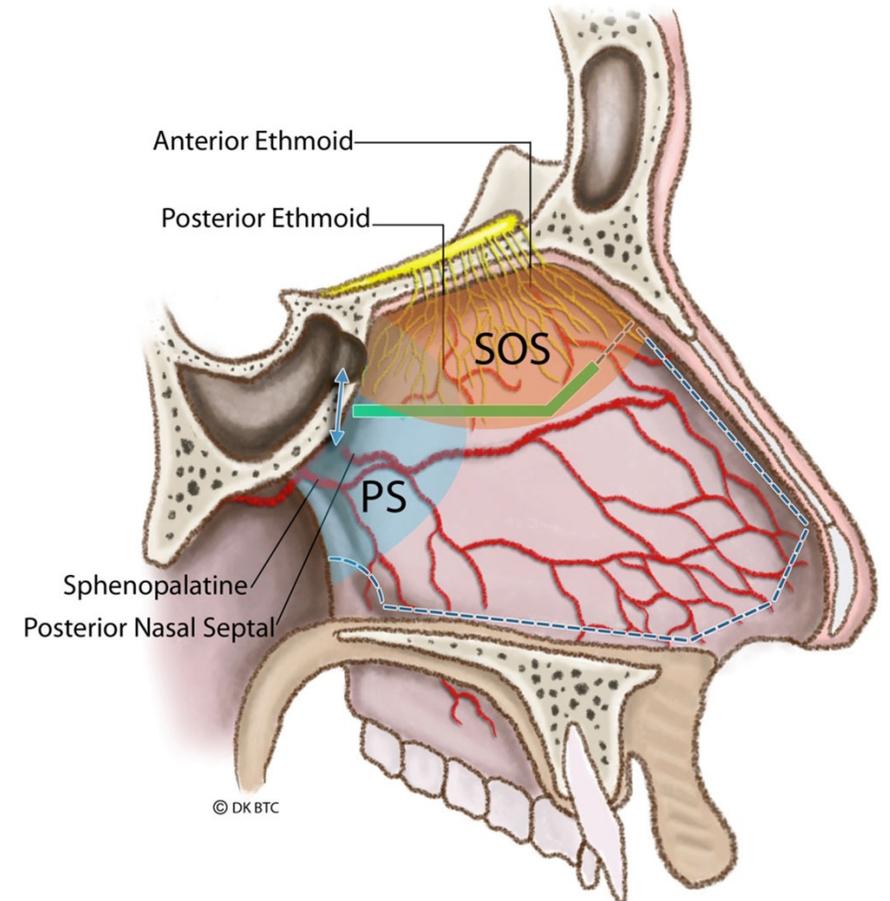
Expanded Approach - Transtubercular





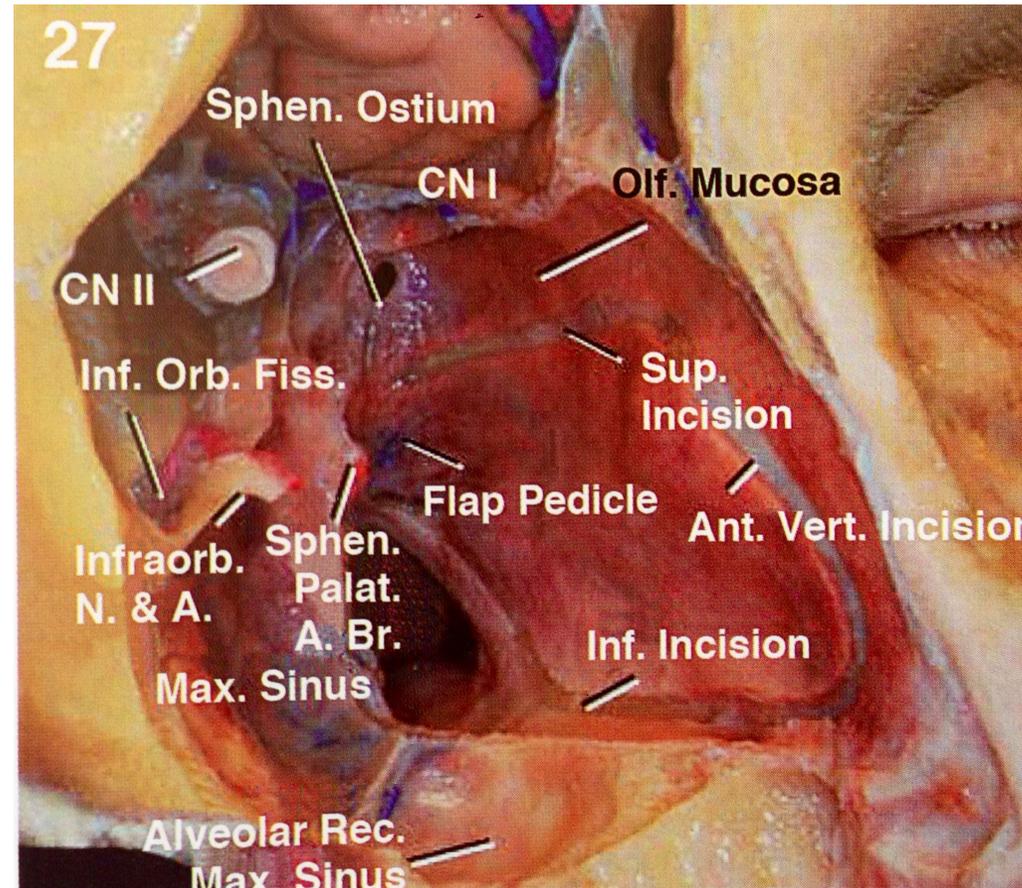
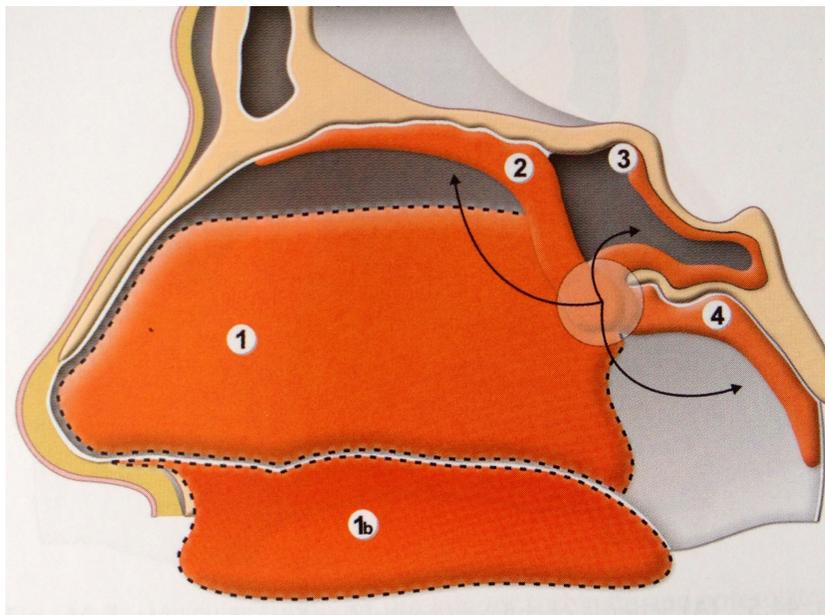
Skull Base Repair

- High rate of post-operative CSF leak (>30%) was a limiting factor for endoscopic skull base.
- With the development of the nasoseptal flap, the CSF leak rate has significantly dropped (<3%).
- Repair paradigm:
 - Dural repair: fascial lata button graft, gasket, inlay duragen + onlay dural substitute
 - Nasoseptal flap.



Nasoseptal Flap

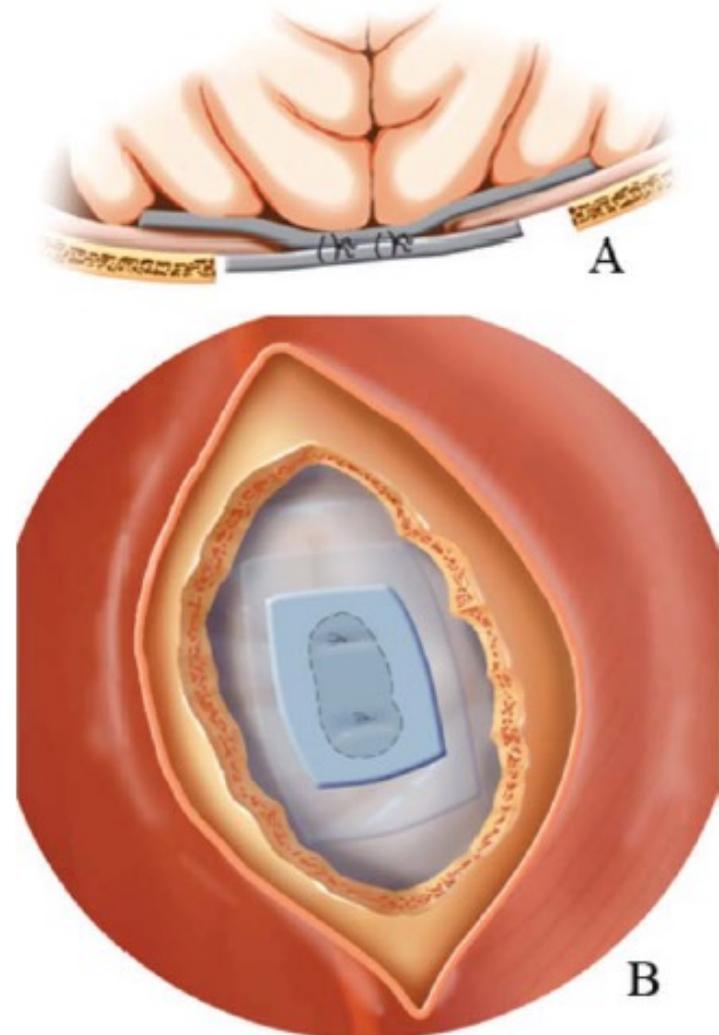
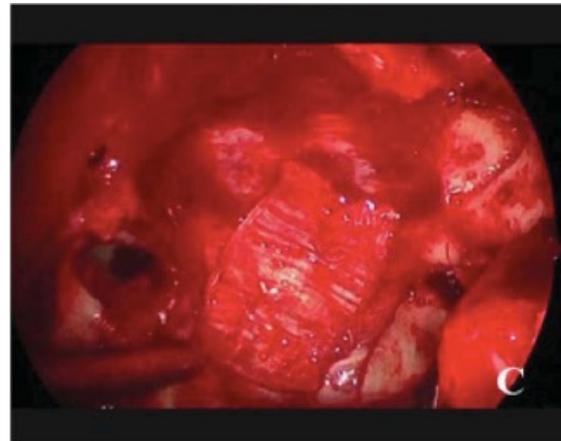
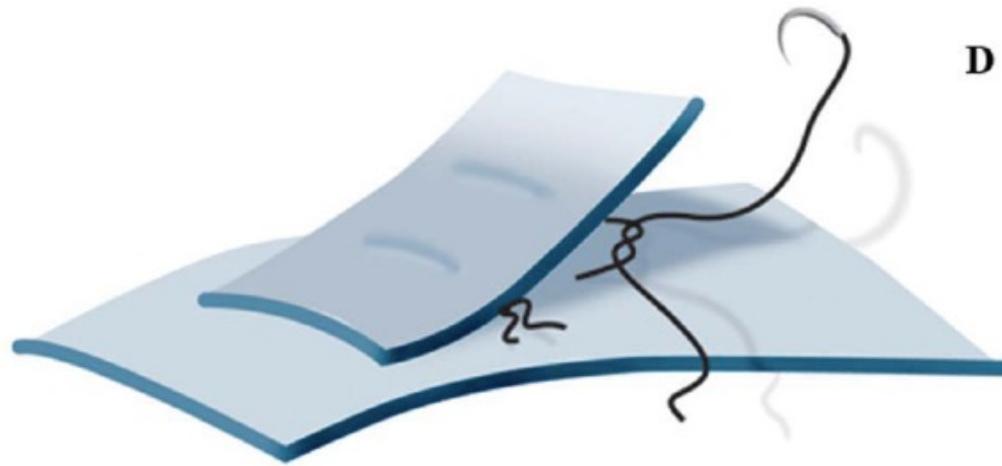
- Pedicled mucosal flap
- Repair of high flow CSF leaks
- Expanded approaches



Ability to raise nasoseptal flap permits repair of large cranial base defects w/o craniotomy

Endoscopic Repair of High-Flow Cranial Base Defects Using a Bilayer Button

Adam J. Luginbuhl, MD; Peter G. Campbell, MD; James Evans, MD; Marc Rosen, MD





Gasket Repair

TJUH Experience

The learning curve in endoscopic endonasal resection of craniopharyngiomas

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- 43 pts from 2005-2015
- Comparison of early and late cohorts to determine the learning curve of endoscopic approaches

TABLE 3. Surgical outcomes

Surgical Outcome	Total	Early (n = 20)	Late (n = 23)	p Value
Histological subtype*				
Adamantinomatous	29 (67%)	14 (74%)	15 (65%)	0.739
Papillary	13 (30%)	5 (26%)	8 (35%)	—
Extent of resection†				
STR	11 (26%)	8 (40%)	3 (13%)	0.078
NTR (>95%)	13 (30%)	8 (40%)	5 (22%)	0.193
GTR	19 (44%)	4 (20%)	15 (65%)	0.005
Complications				
New hydrocephalus	5 (12%)	4 (20%)	1 (4%)	0.167
Meningitis	2 (5%)	2 (10%)	0	0.210
CSF leak	9 (21%)	8 (40%)	1 (4%)	0.007
Hyponatremia	1 (2%)	0	1 (4%)	—
Seizure	1 (2%)	0	1 (4%)	—
Medical complications‡	6 (14%)	4 (20%)	2 (9%)	0.393
Rhinological complications§	8 (19%)	4 (20%)	4 (17%)	—
Carotid artery injury	1 (2%)	1 (5%)	0	0.465
Stroke	1 (2%)	1 (5%)	0	0.465
Median length of stay (days)	6	6	6	—
Discharge status¶				
Home	30 (83%)	9 (64%)	21 (95%)	0.024
Rehabilitation or nursing facility	6 (17%)	5 (36%)	1 (5%)	—

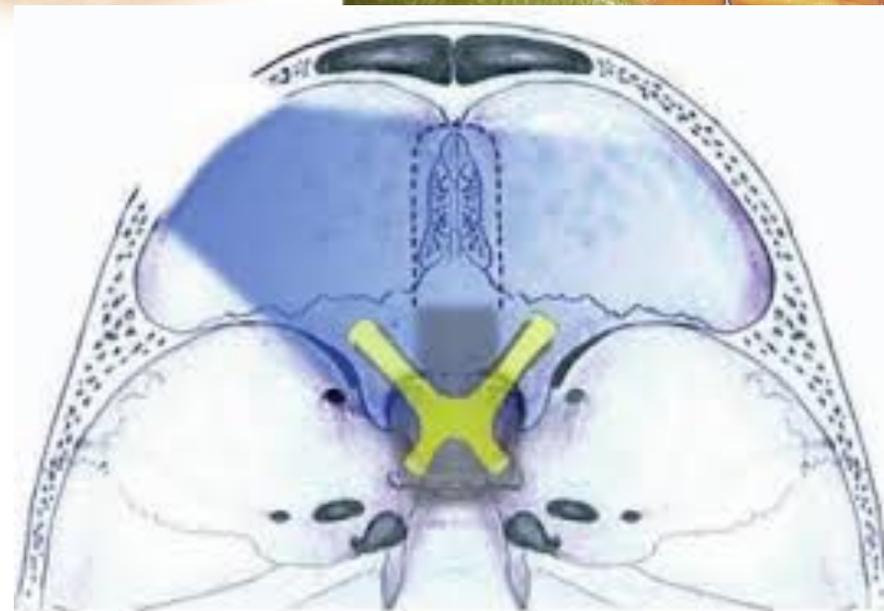
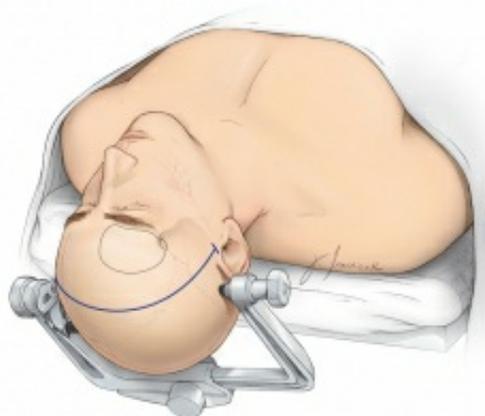
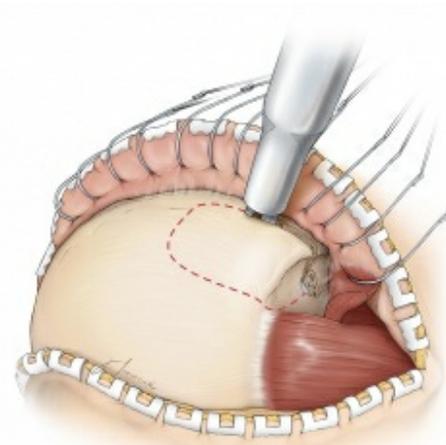
TABLE 4. Postoperative visual outcome*

Visual Outcome	Total	Early (n = 20)	Late (n = 23)	p Value
Improved	31 (84%)	14 (88%)	17 (81%)	0.680
Normalized	9 (24%)	6 (38%)	3 (14%)	—
Stable	5 (14%)	1 (6%)	4 (19%)	—
Worse	1 (3%)	1 (6%)	0	—



“Eyebrow” Supra-Orbital Craniotomy

- “Sweet-spot” parasellar lesions



Conclusion

- Endoscopic skull base surgery offers a less invasive approach for safe resection of pituitary region tumors.
- Requires a multidisciplinary team.
- Steep learning curve therefore fellowship training is essential.
- Its limitations should be acknowledged.
- Use of endoscope can lead to less invasive open transcranial approaches by offering a wider angle of view.

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