

Curriculum Vitae

Rahadian Indarto Susilo, dr SpBS

- Fak Kedokteran Universitas Airlangga 1996-2002
- PPDS Bedah Saraf Universitas Airlangga Surabaya 2003-2009
- Skull Base Fellow Program Keio University Tokyo Japan, 2009
- WFNS Grade A: Skull Base Fellowship Program Osaka City University, Japan. 2011
- Neurooncology Division, Neurosurgery Department Airlangga University- Dr. Sutomo General Hospital, Surabaya, 2009 - now

BRAIN TUMOR UPDATE 2018

Tumor Otak

TEHNOLOGI & PENANGANAN TERKINI

**Rahadian Indarto Susilo,
Irwan Barlian Imadoel Haq, Joni Wahyuhadi**

Divion Of Neuro-Oncology & Skull Base Neurosurgery

Dept. of Neurosurgery, Airlangga Univ. Faculty Of Medicine / Dr. Sutomo General Hospital

SURABAYA INDONESIA



**BRAIN TUMOR
& PITUITARY
DISORDER
CENTER**

RS PREMIER SURABAYA

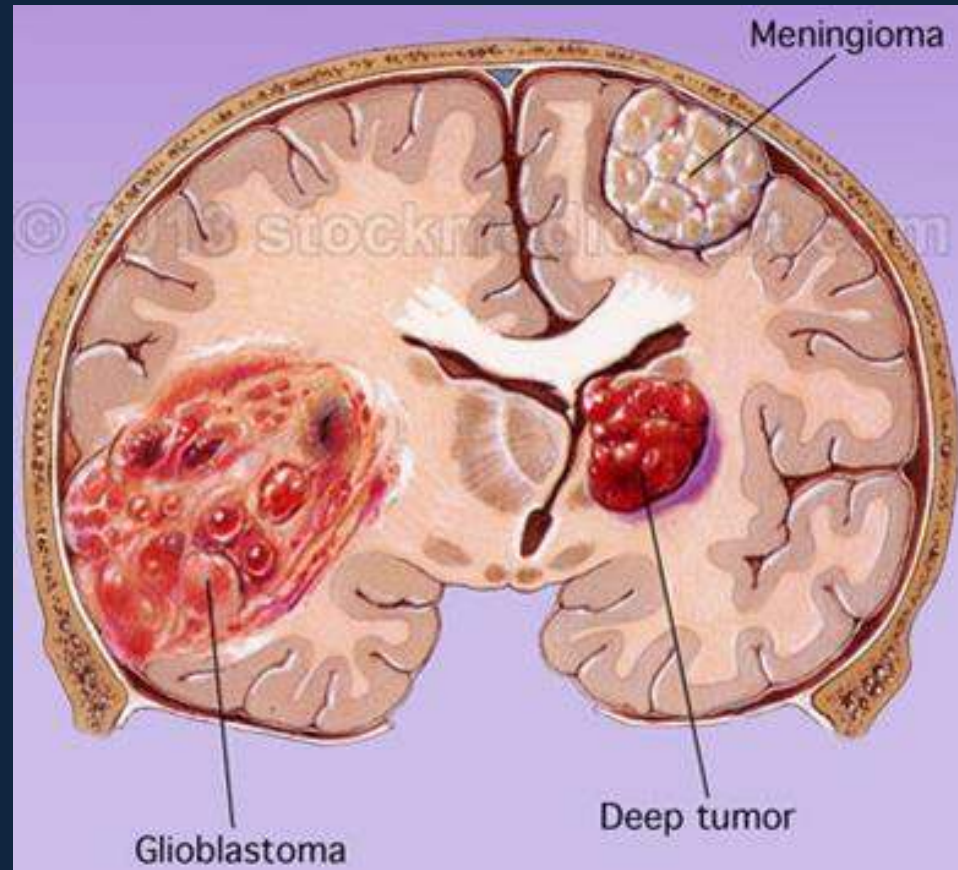
www.BrainTumorIndonesia.com



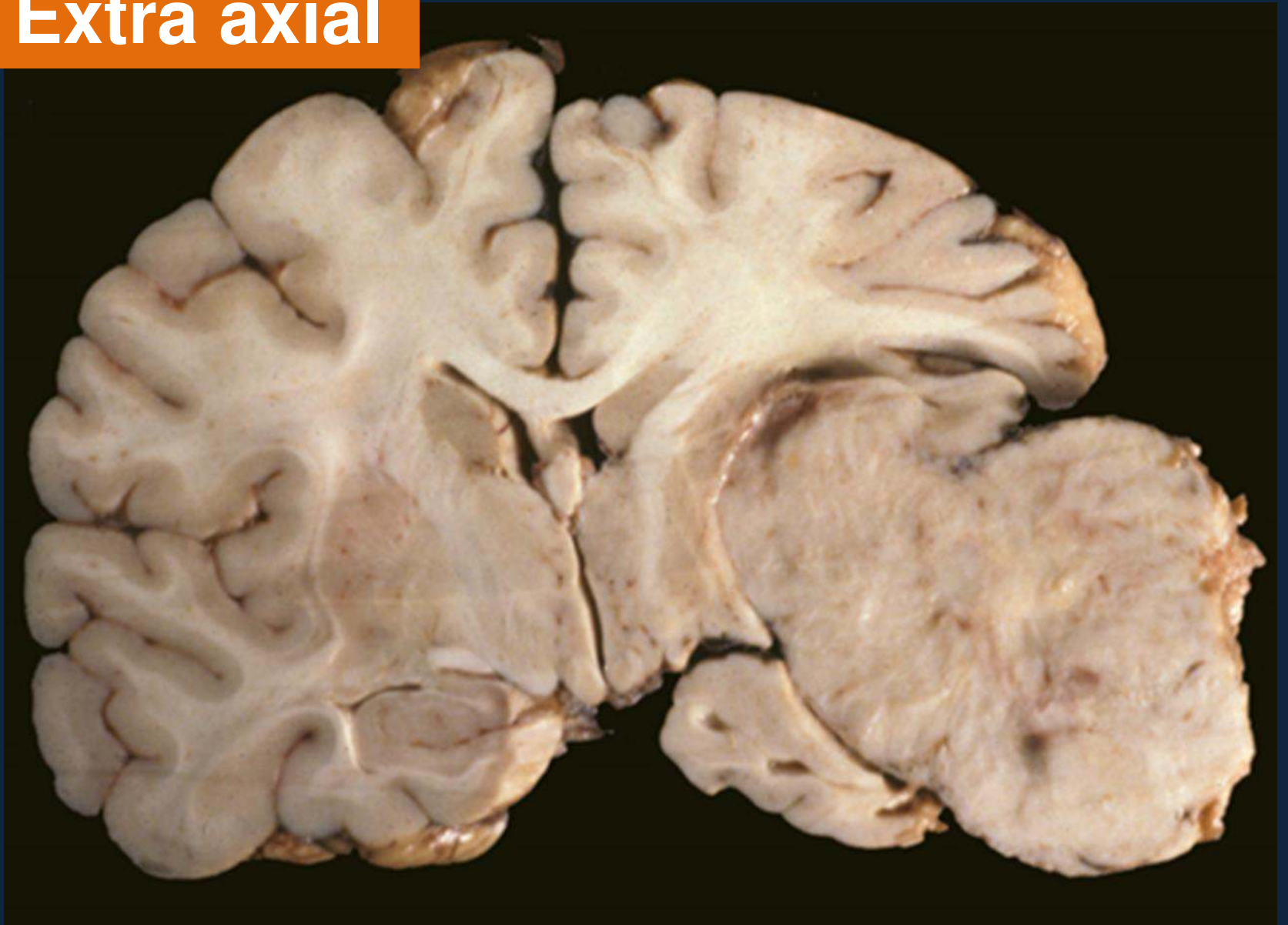
RUMAH SAKIT UMUM DAERAH
Dr. SOETOMO

Tumor otak

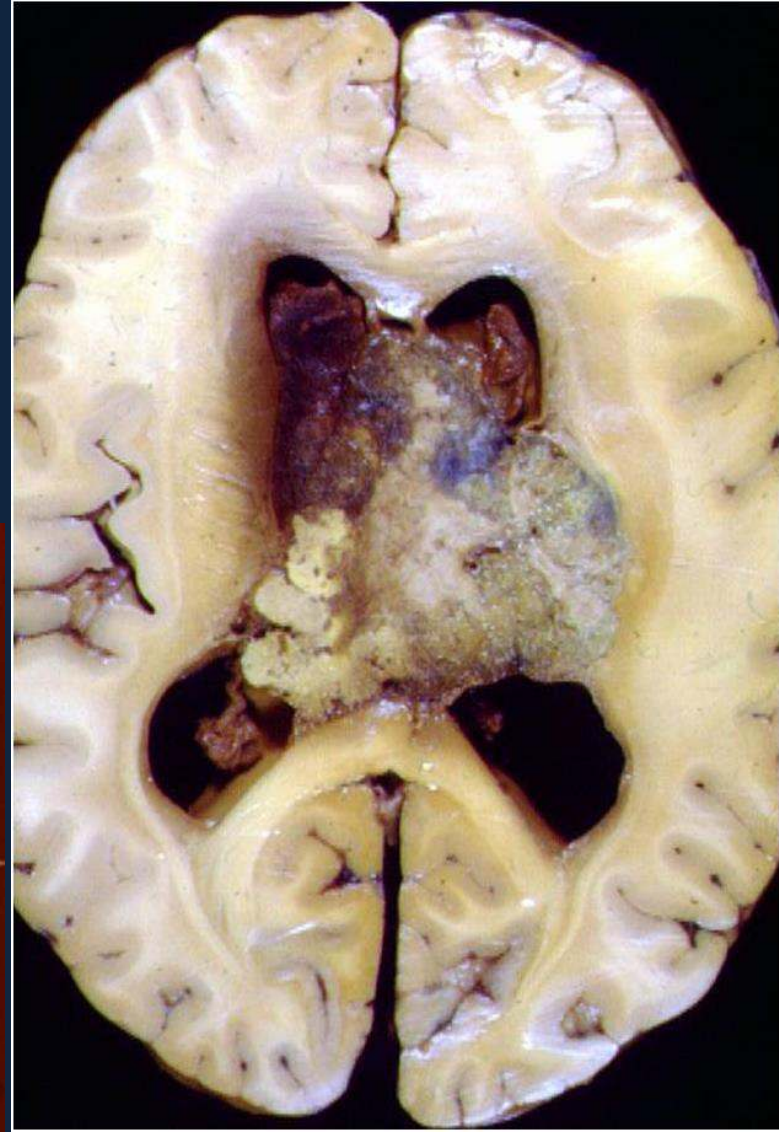
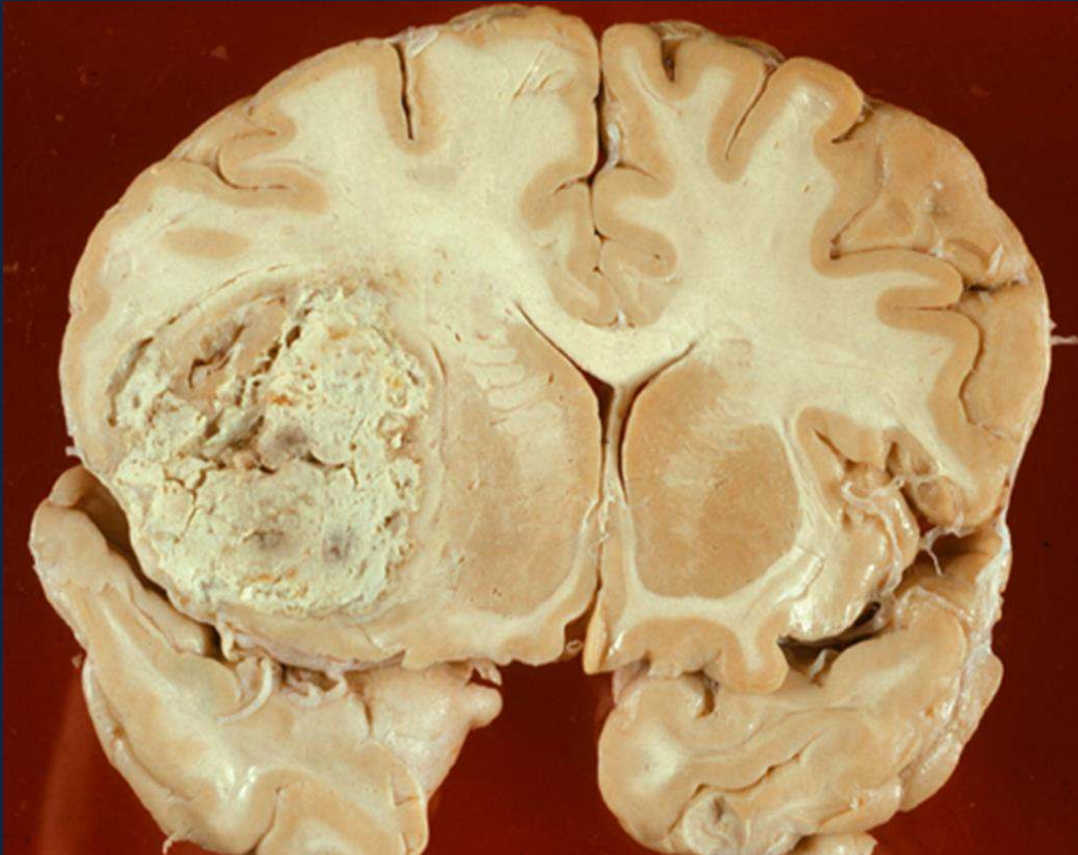
Pertumbuhan jaringan abnormal yang tidak terkendali di dalam otak



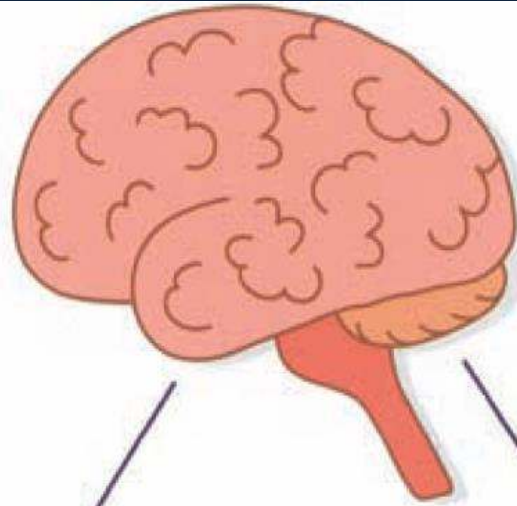
Extra axial



INTRA AXIAL



Clinical features



Clinical Features

- Low mitotic index
- Diffuse invasion
- High rate of transformation

**Low grade
Astrocytoma**

5-10 years

**Primary
Glioblastoma**

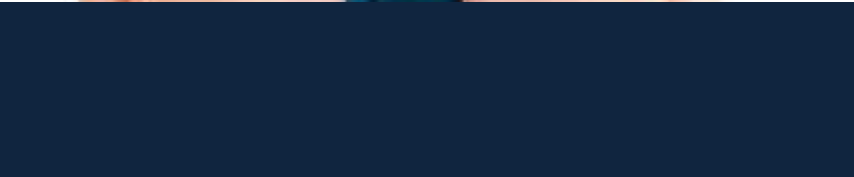
**Secondary
Glioblastoma**

Clinical Features

- Rapid proliferation
- Diffuse invasion
- Angiogenesis
- Cellular necrosis

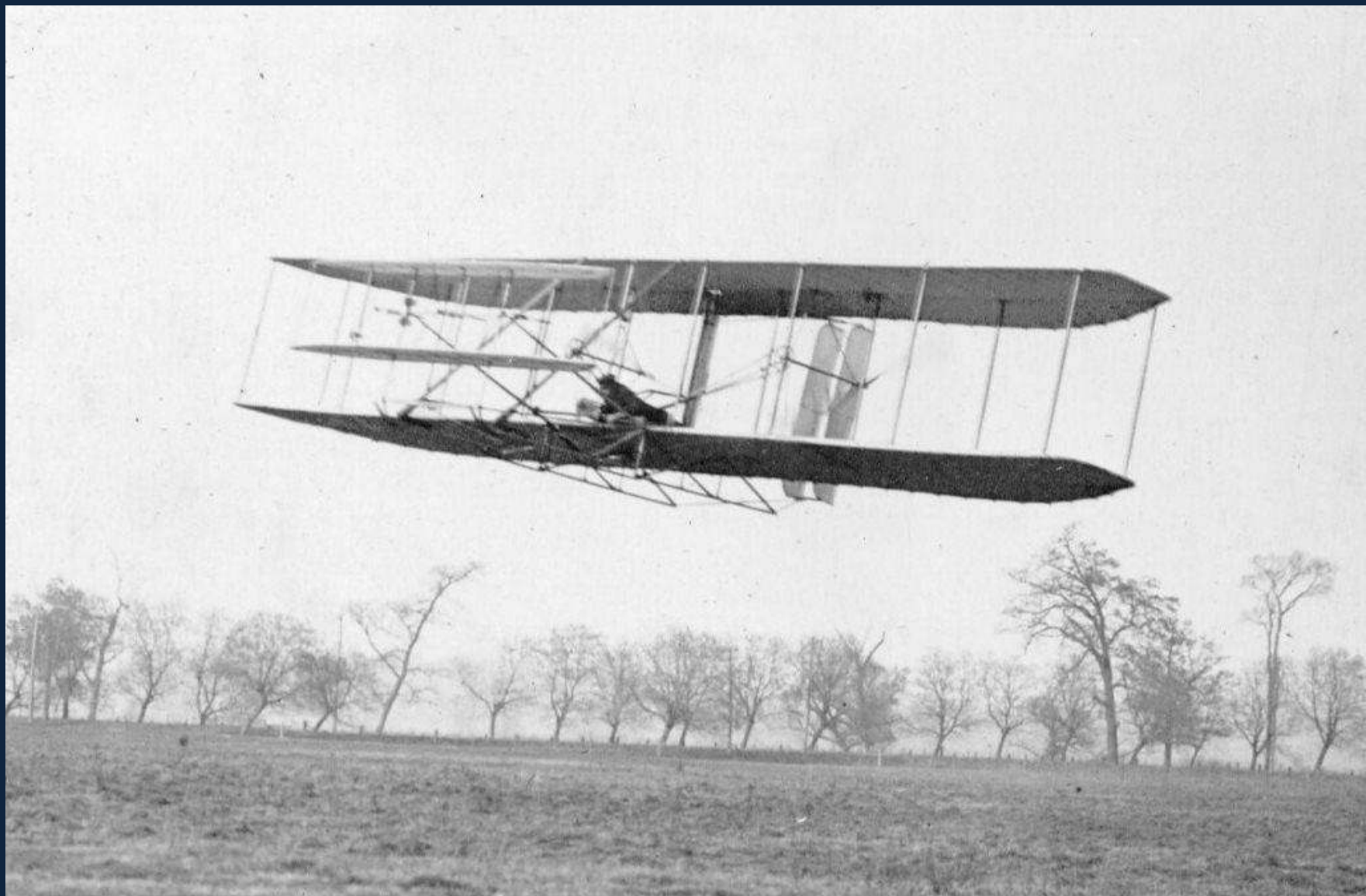
Terapi Tumor Otak

OPERASI

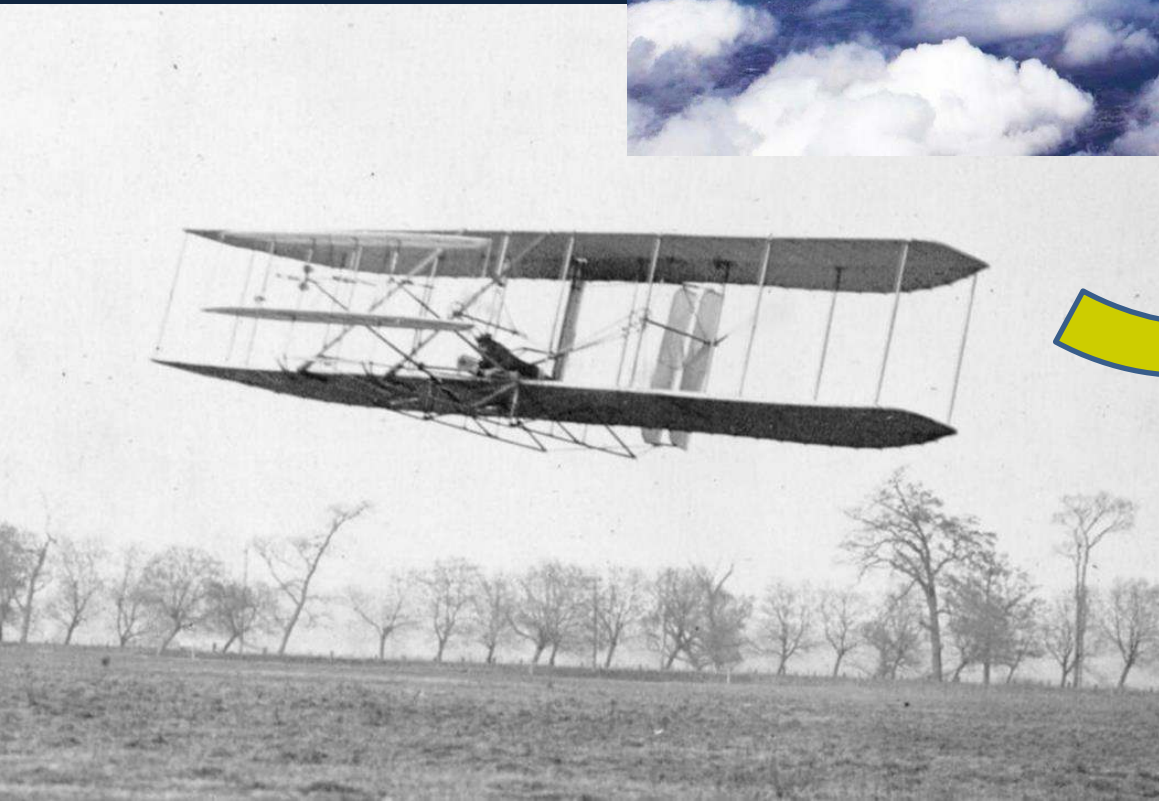


Apakah AMAN ???









Terapi Tumor Otak

OPERASI

**KEEP YOUR
BRAIN SAFE**

**WHAT DO YOU
MEAN**

???





*Luck is what
happens when
preparation meets
opportunity.*

- Seneca

PREPARATION

Good Preparation → GREAT Preparation

GOOD is the enemy of

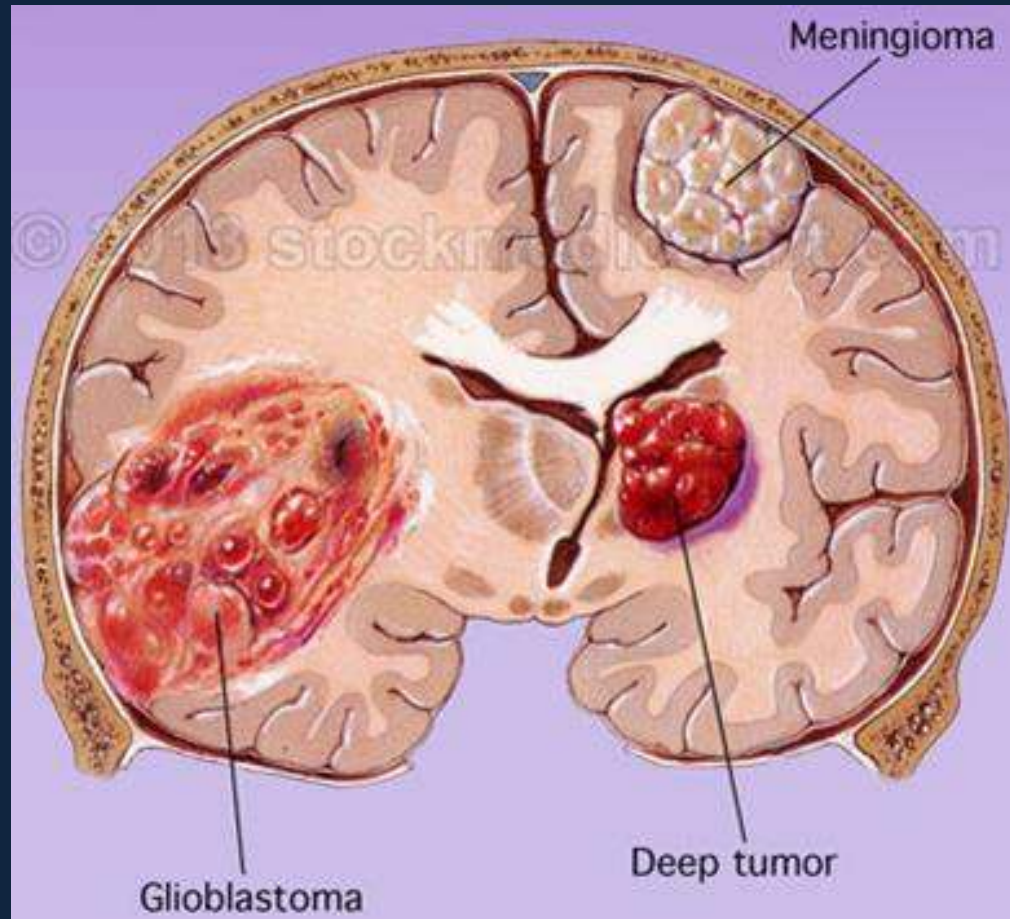


PREPARATION

Good Preparation → GREAT Preparation

GREAT NEURO-IMAGING

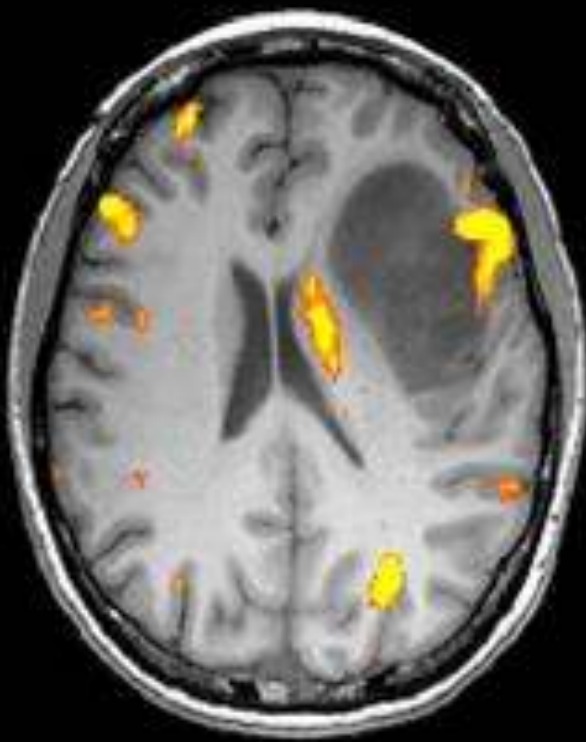
Bagaimana membedakan ????



jaringan tumor v/s jaringan otak normal

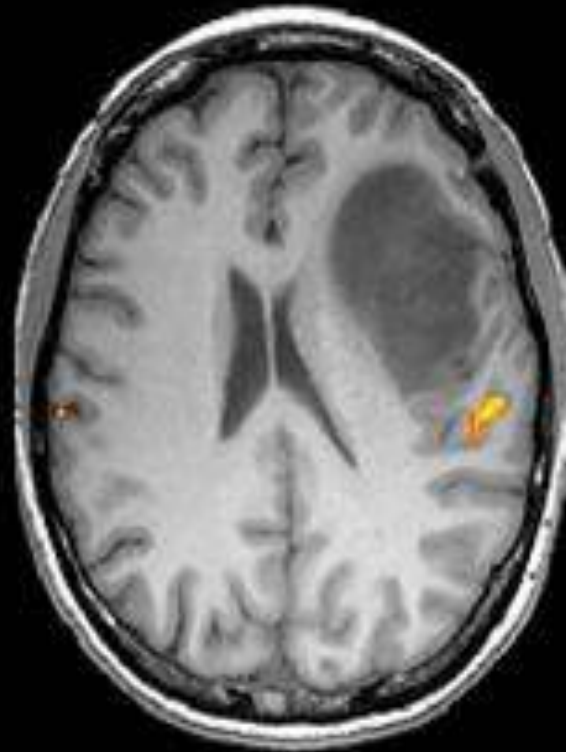
Functional MRI

pada TUMOR OTAK



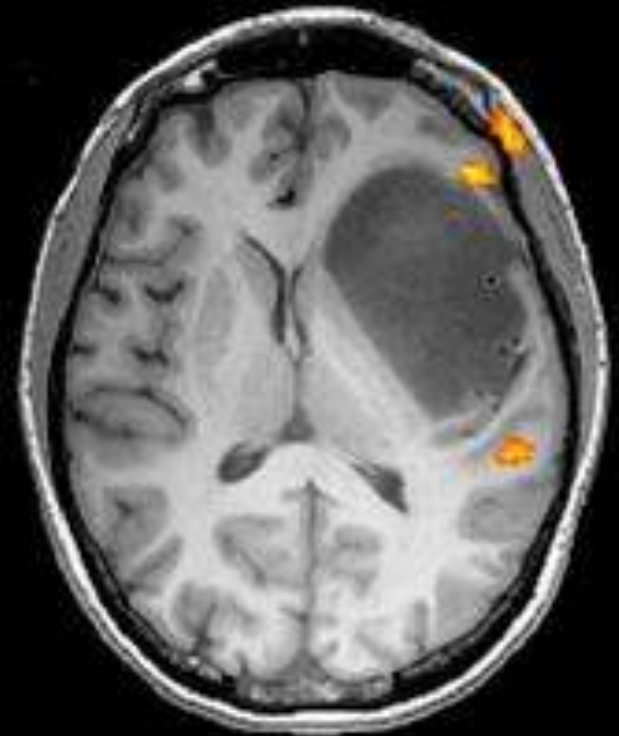
speech

AREA BICARA



finger tap

AREA MOTORIK



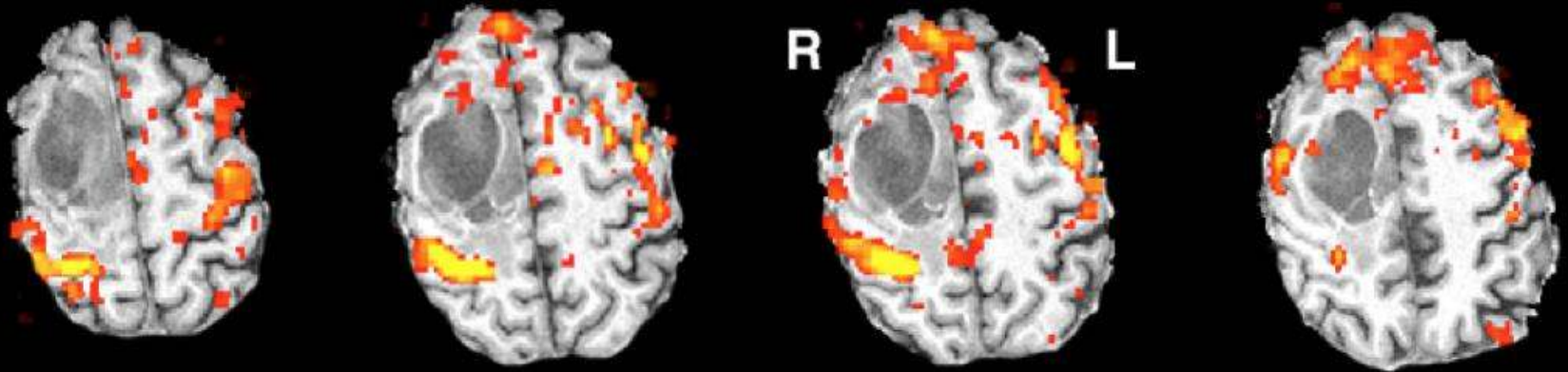
listening

AREA BICARA

Functional MRI pada TUMOR OTAK

IDENTIFIKASI AREA MOTORIK

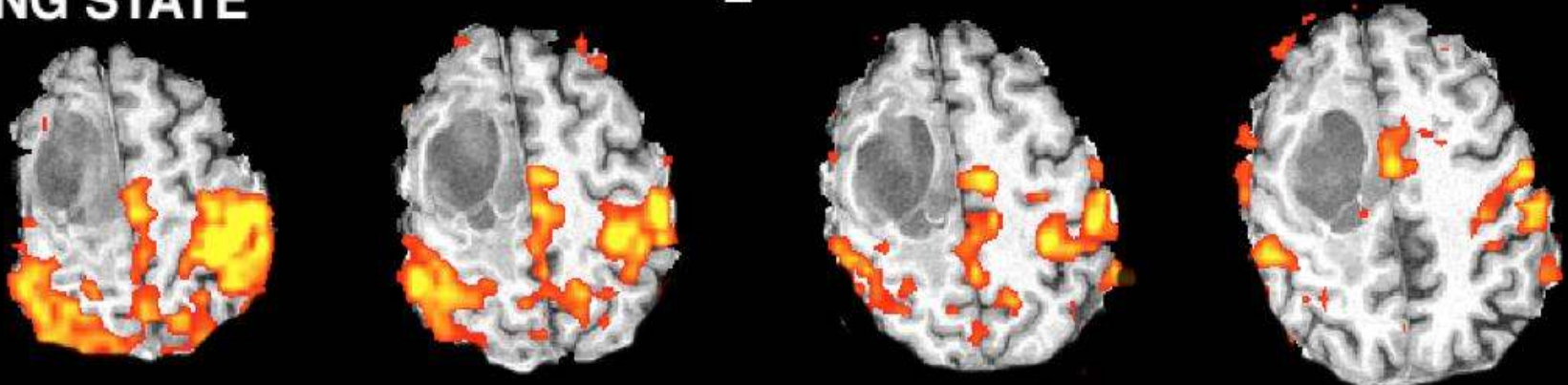
LEFT HAND MOVEMENT



3.1 7.0

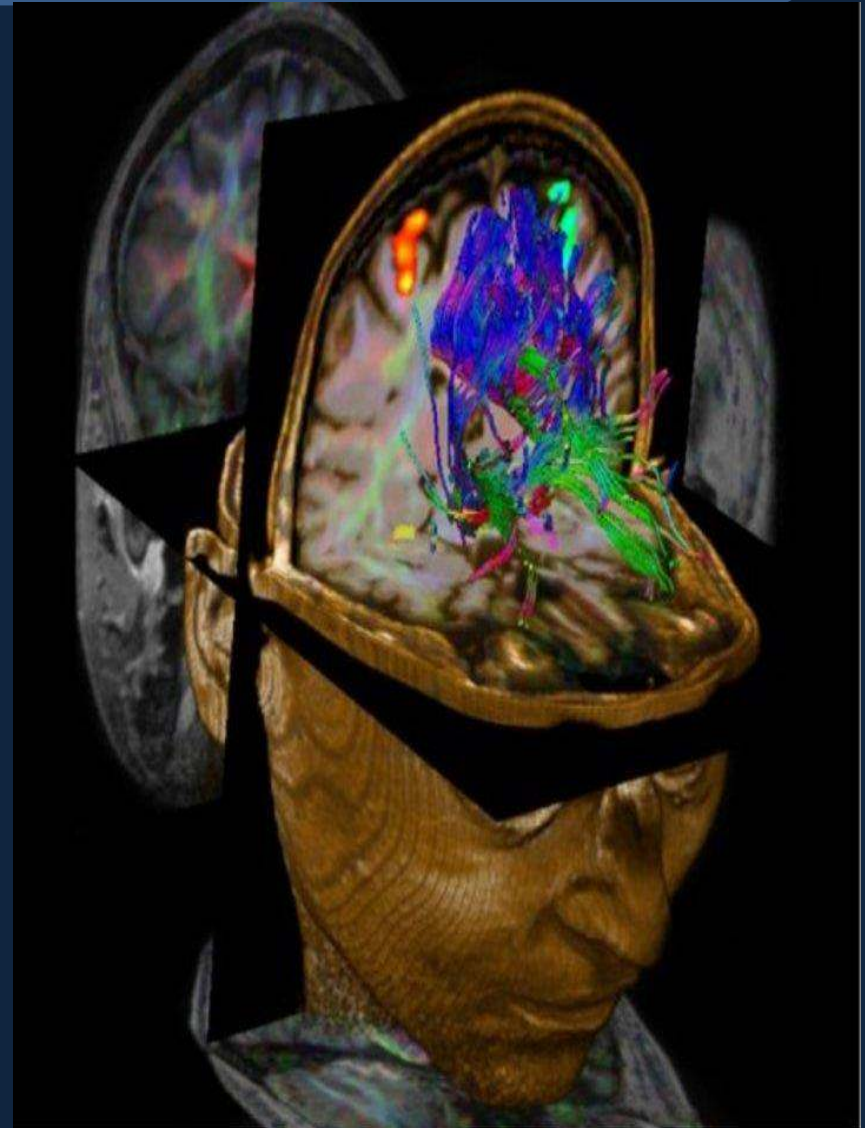
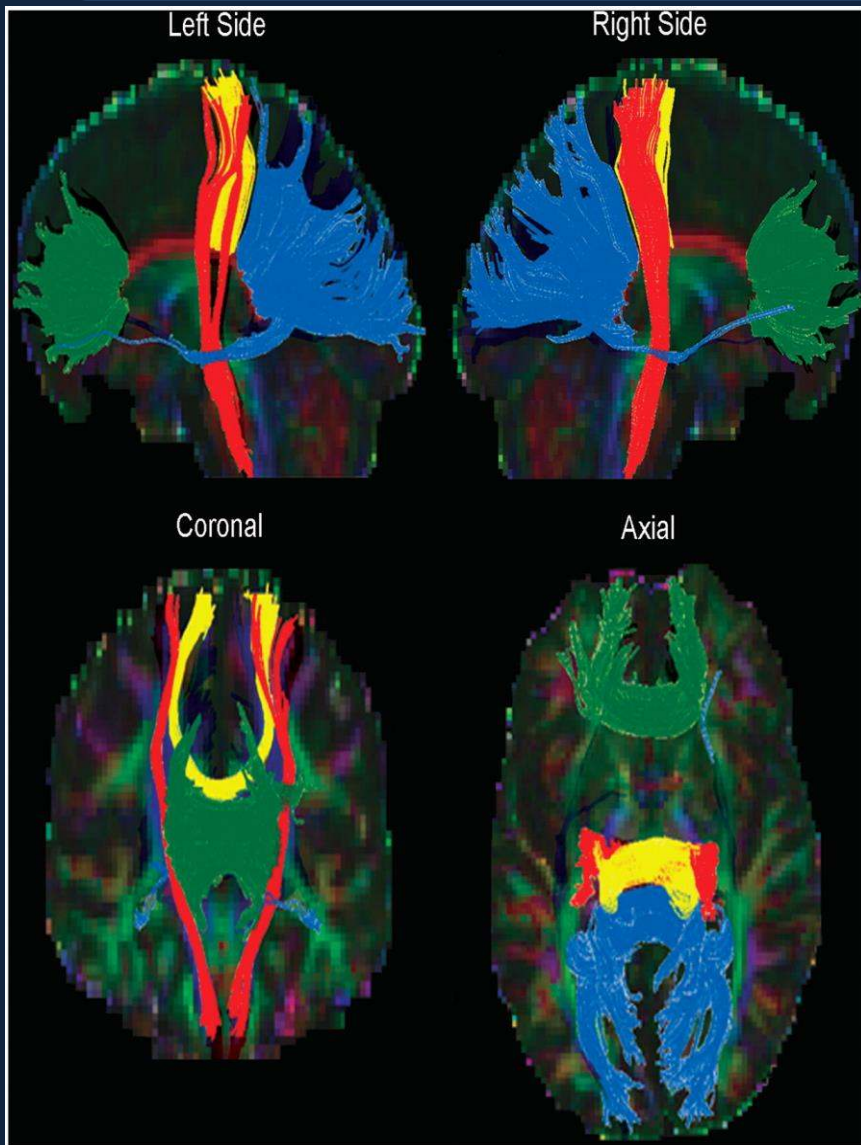
Z

RESTING STATE

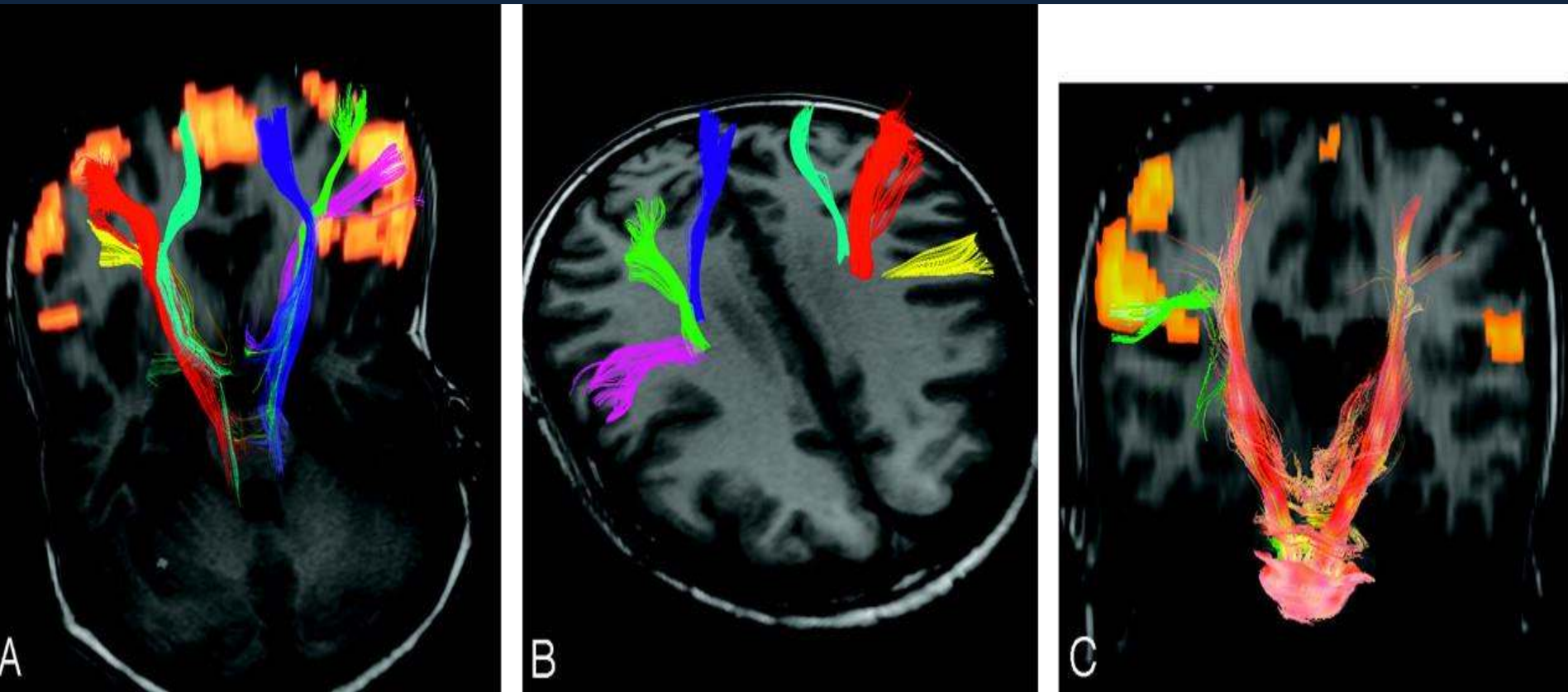


MRI Tractography

IDENTIFIKASI JARAS DI DALAM OTAK

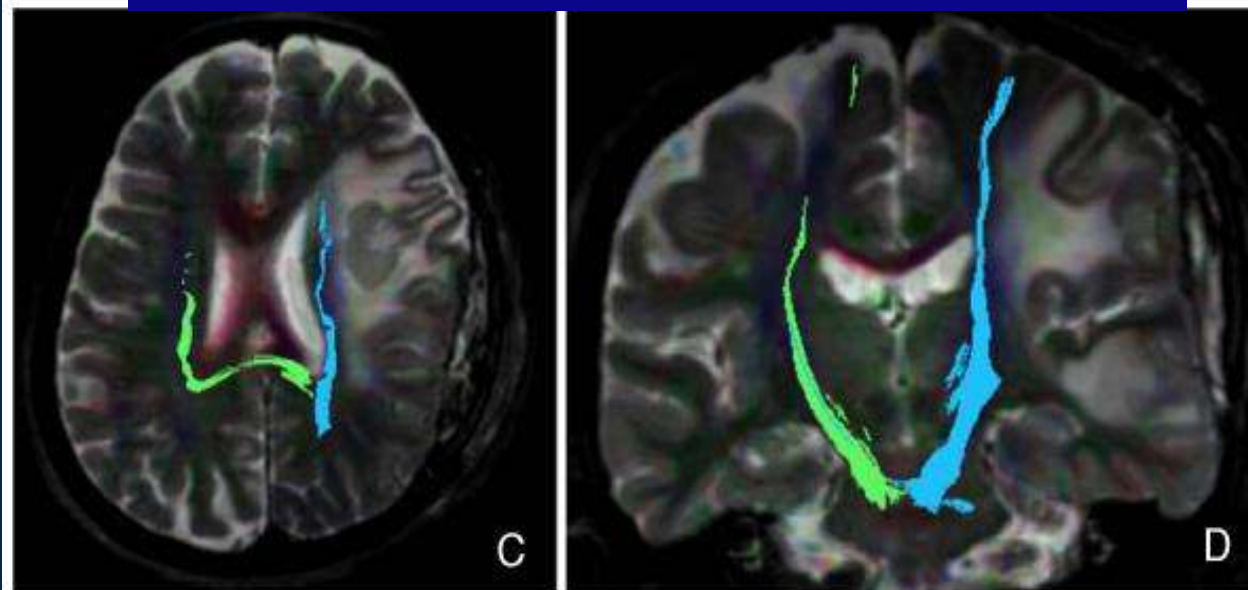
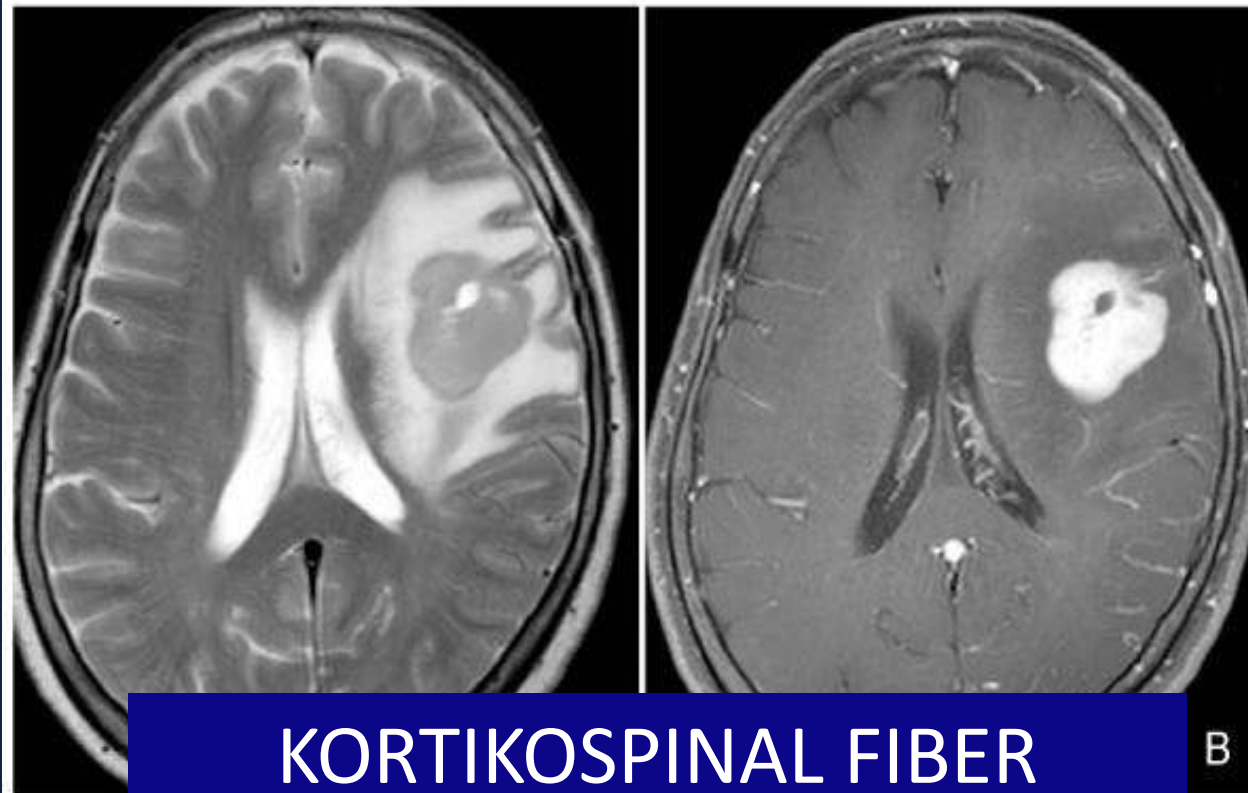


MRI Tractography

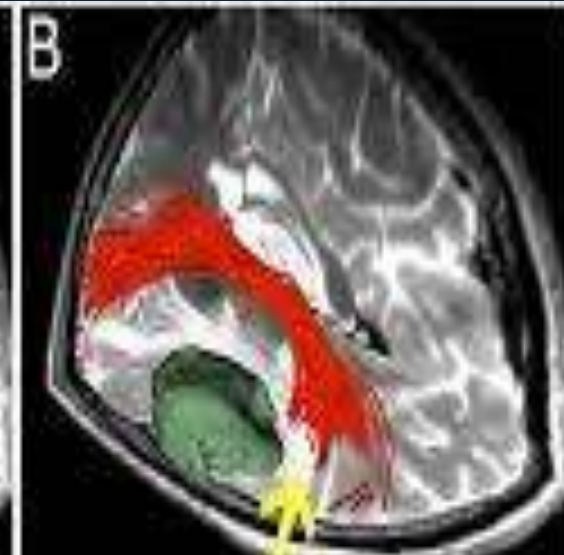
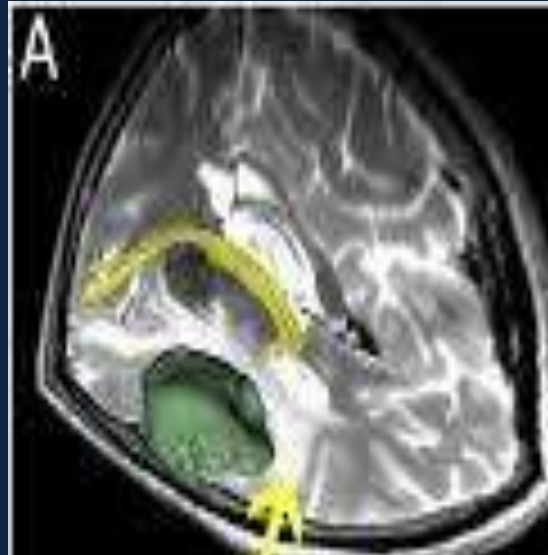
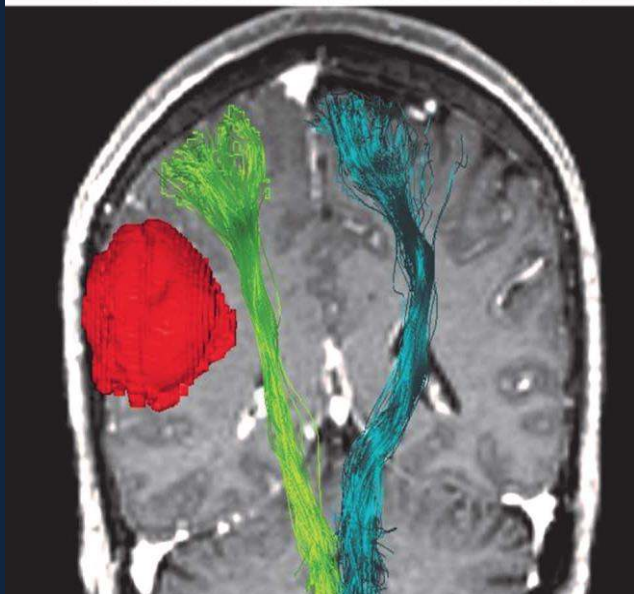
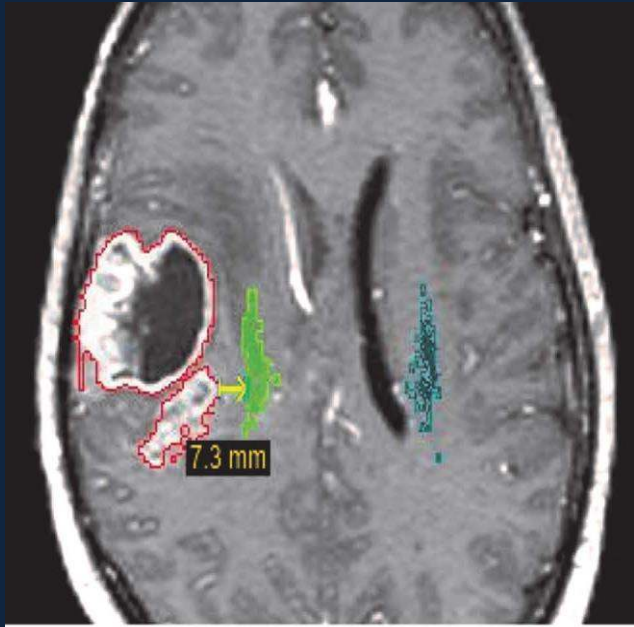


JARAS KORTIKOSPINAL

TRACTOGRAPHY
pada
TUMOR OTAK



MRI Tractography pada TUMOR OTAK



KORTIKOSPINAL FIBER



OPERASI BEDAH SARAF

Prioritas Utama

KUALITAS HIDUP PASIEN

Prioritas UTAMA

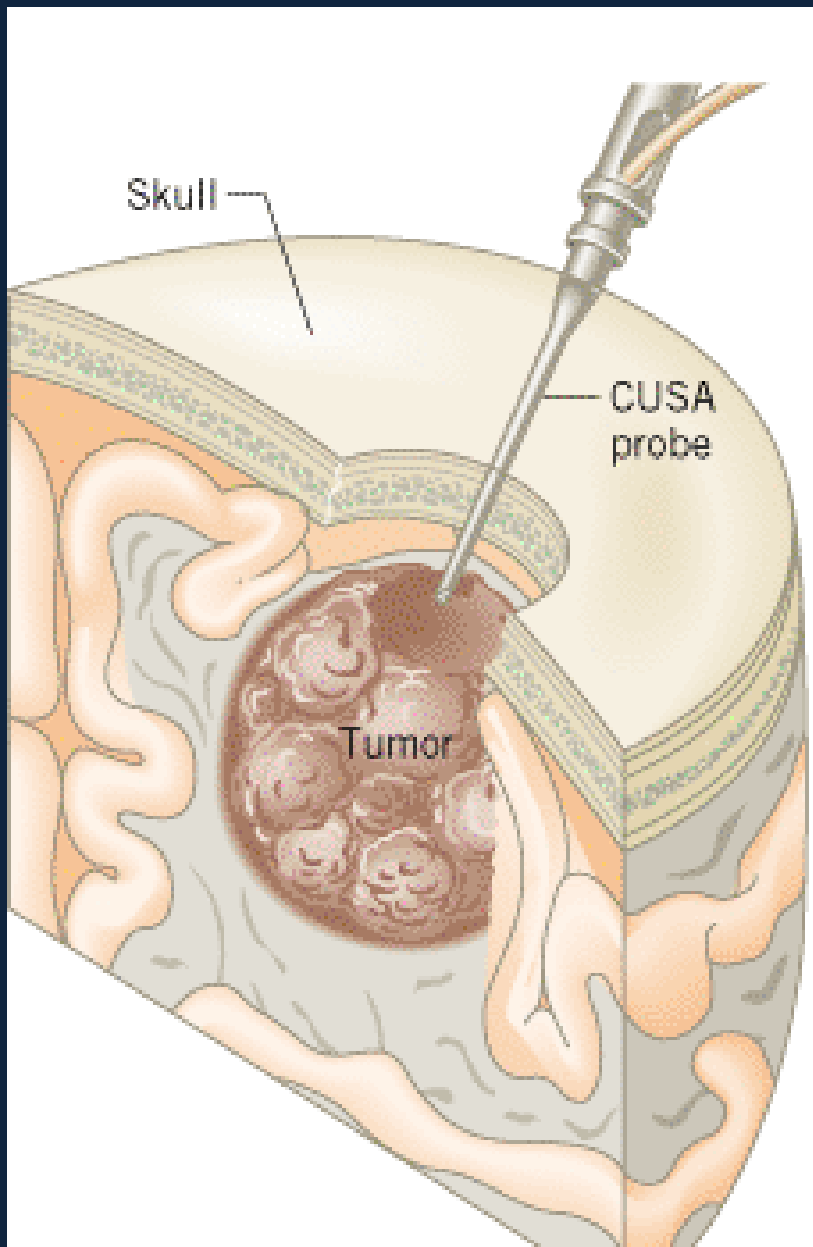
Keamanan Proses Pembedahan

**KEEP YOUR
BRAIN SAFE**

Microneurosurgery

membedakan jaringan otak dan tumor

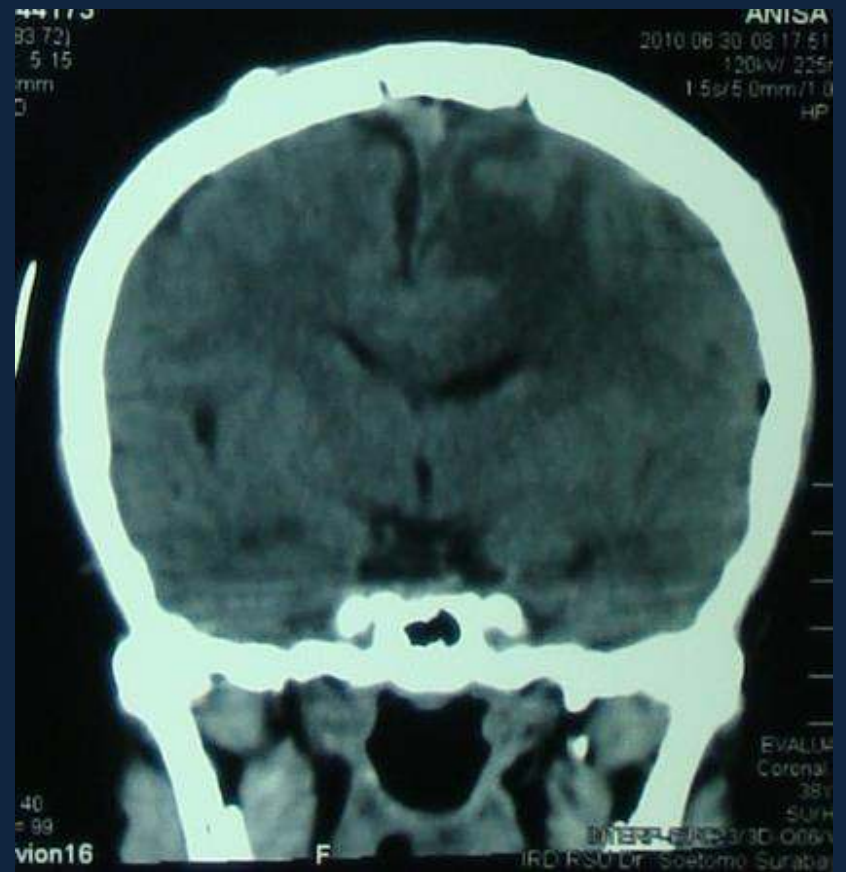
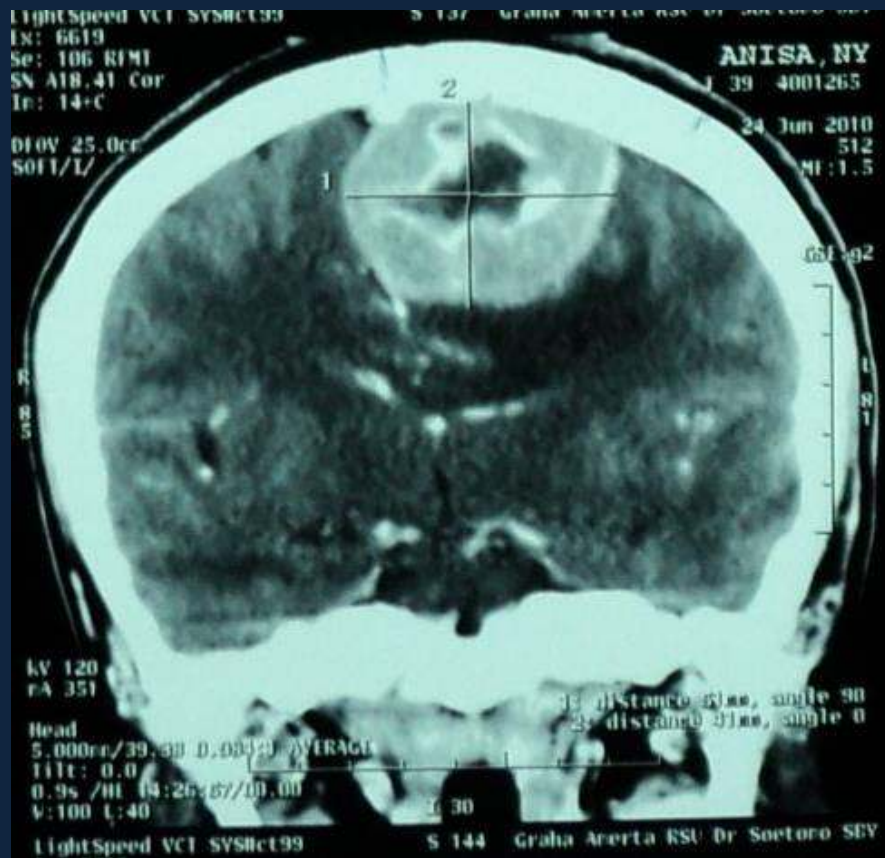




Alat Khusus Penghancur Tumor



cavitron ultrasonic surgical aspirator (CUSA)



Keamanan Proses Pembedahan

Intra-operative monitoring



- Evaluasi fungsi saraf
- Area fungsi bicara
- Saraf wajah
- Saraf perasa
- Saraf penggerak



Keamanan Proses Pembedahan

Intra-operative monitoring

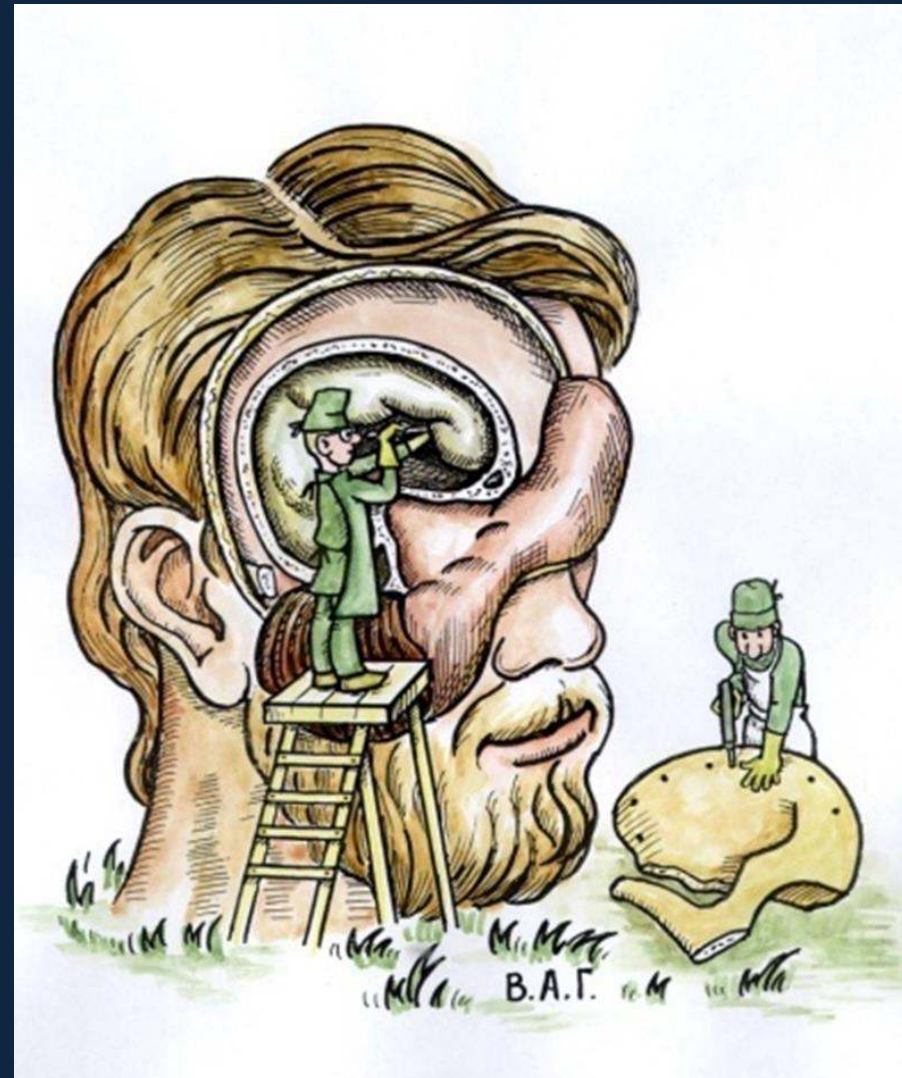
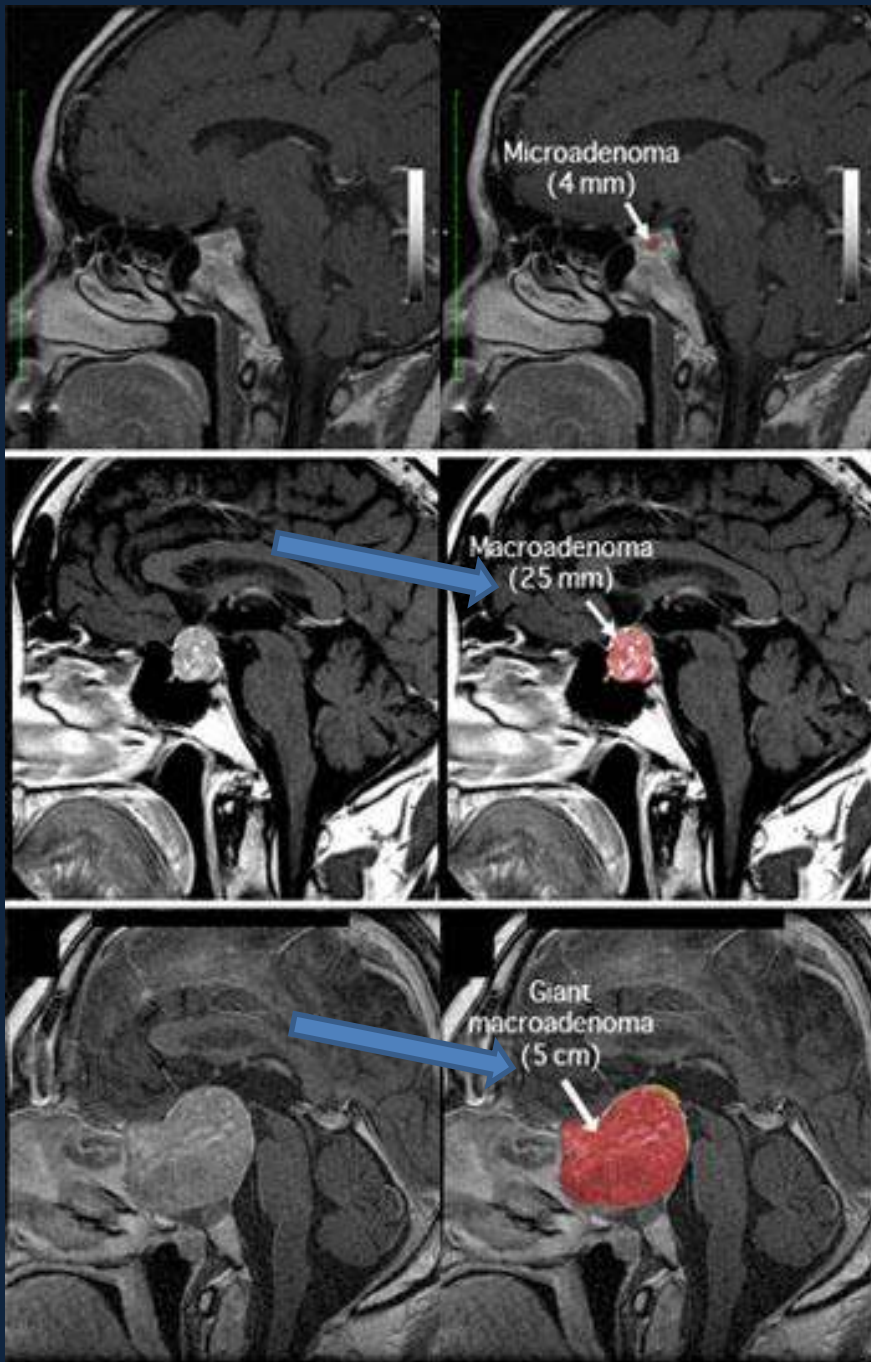


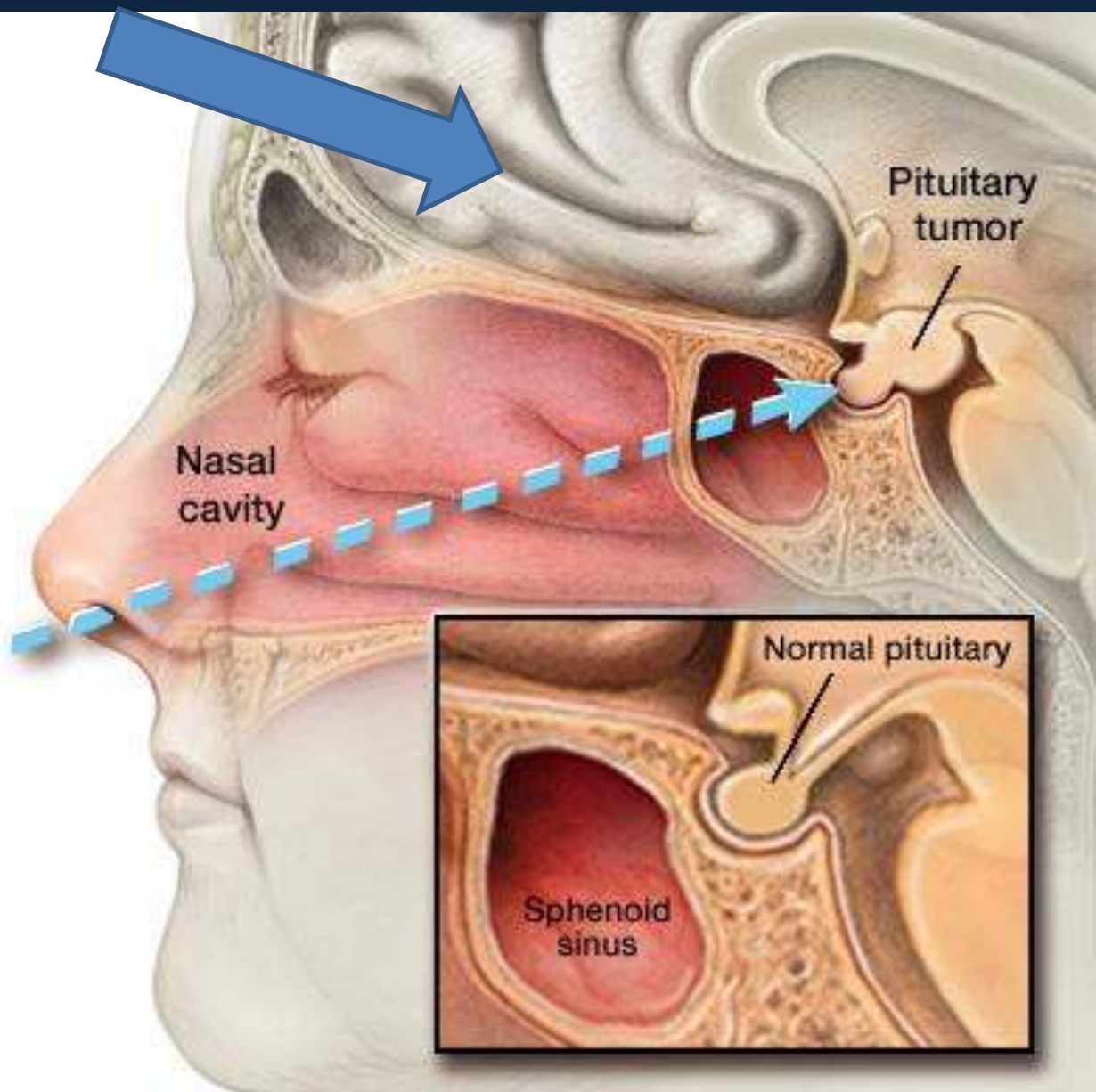
Minimal invasif

Endoscopic

Neurosurgery

Kraniotomi

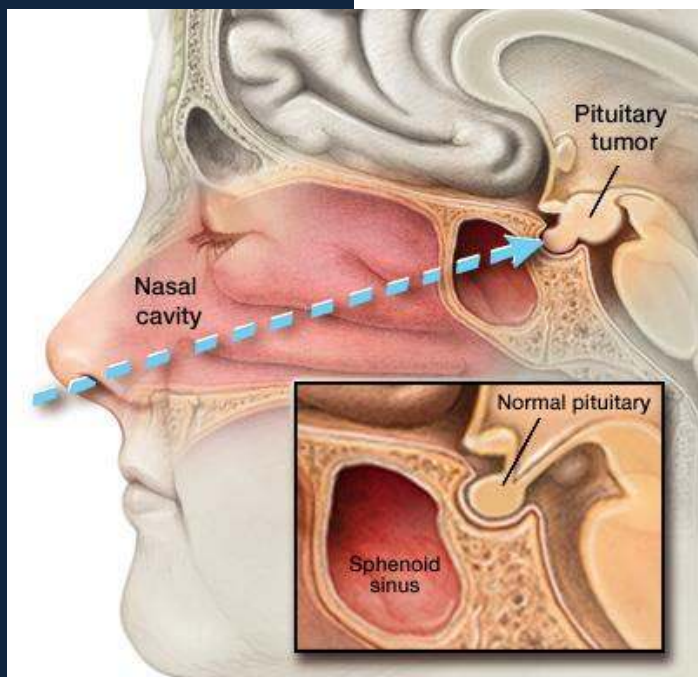




**MINIMAL
INVASIF**

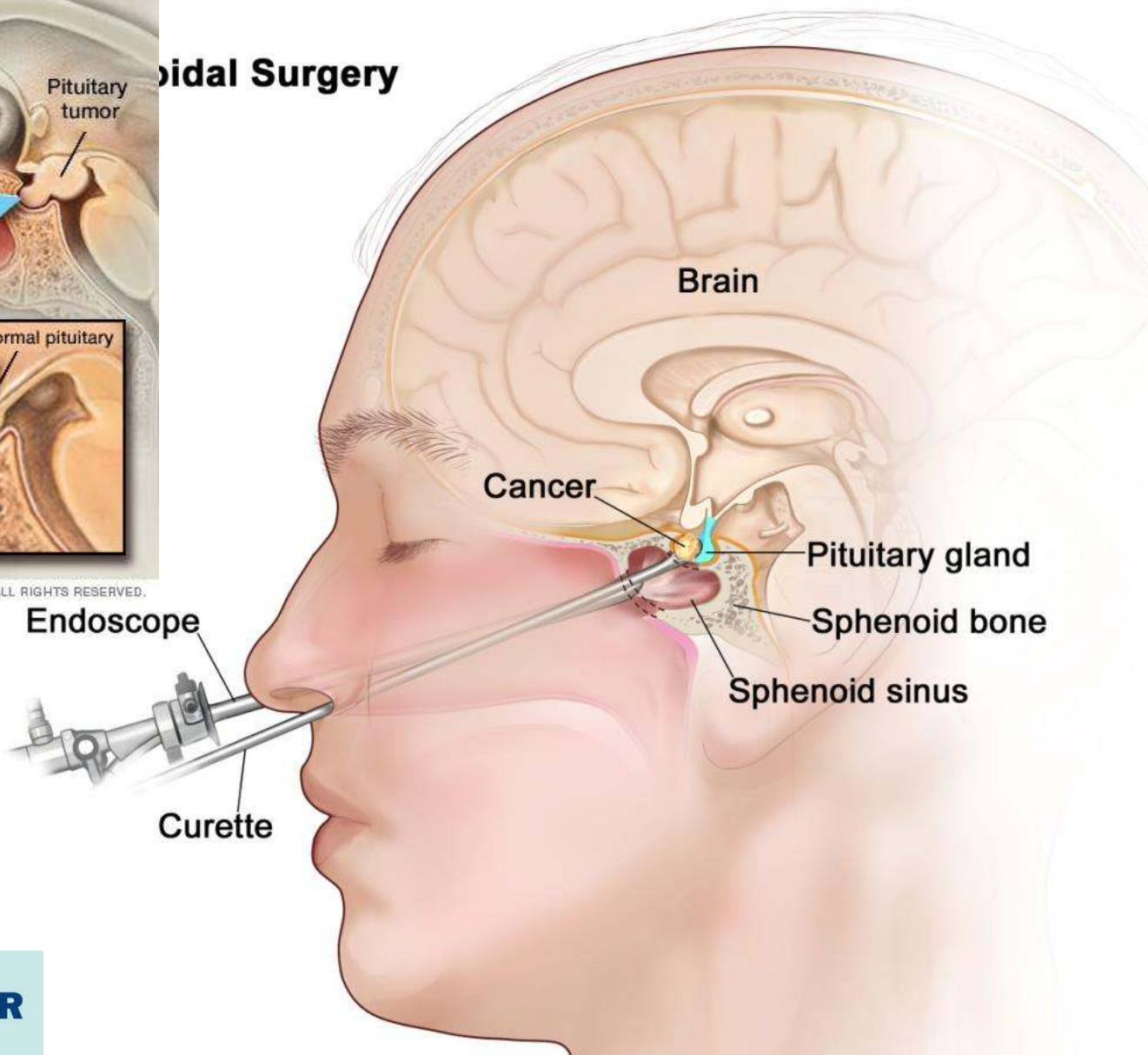
**Endoskopi
Endonasal
neurosurgery**

Endoskopi neurosurgery



Endoscopic Pituitary Surgery

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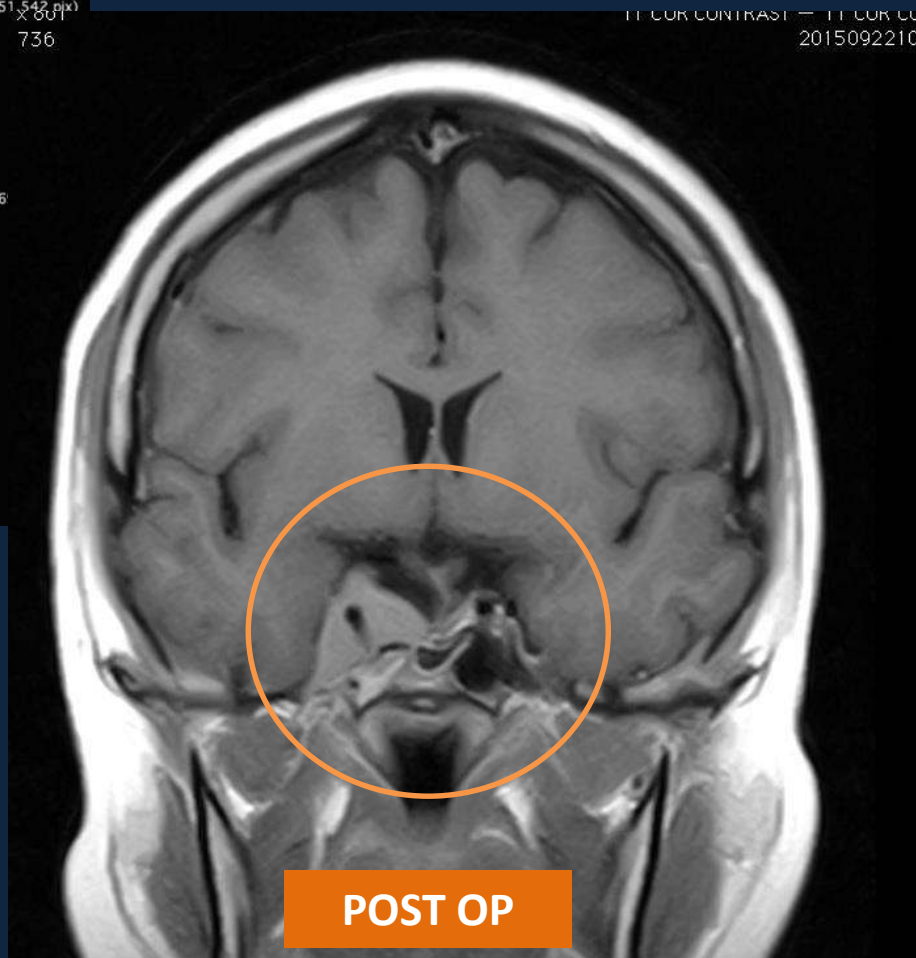
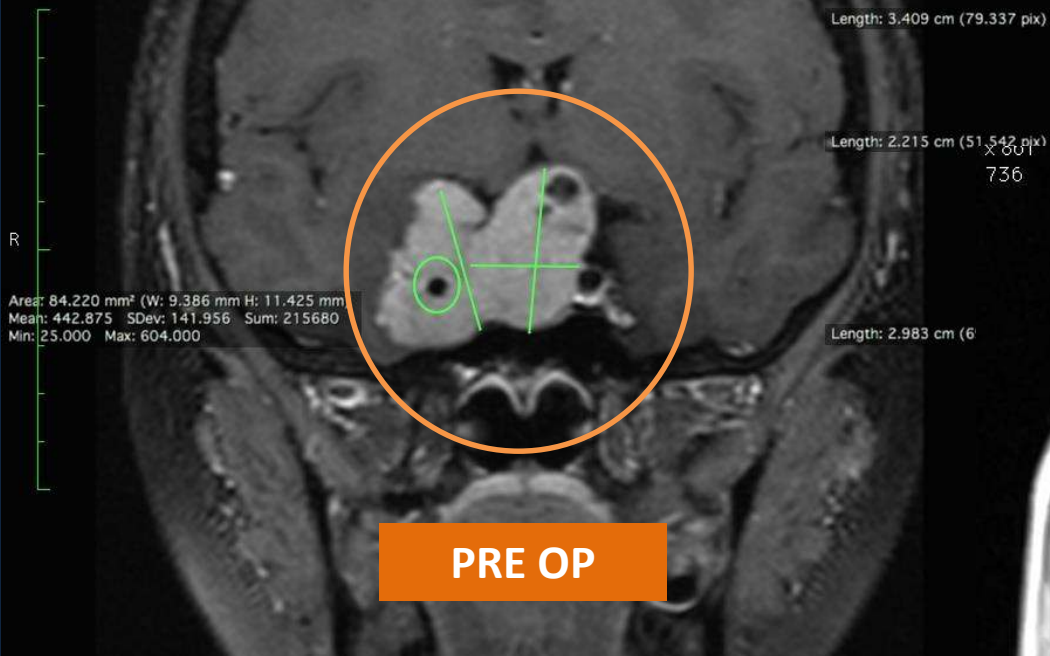
**Pria, 23 tahun,
Gangguan Penglihatan (LAPANG PANDANG)
Gangguan ereksi
Peningkatan Hormon PROLAKTIN**



Image size: 512 x 512
View size: 903 x 861
WL: 478 WW: 976

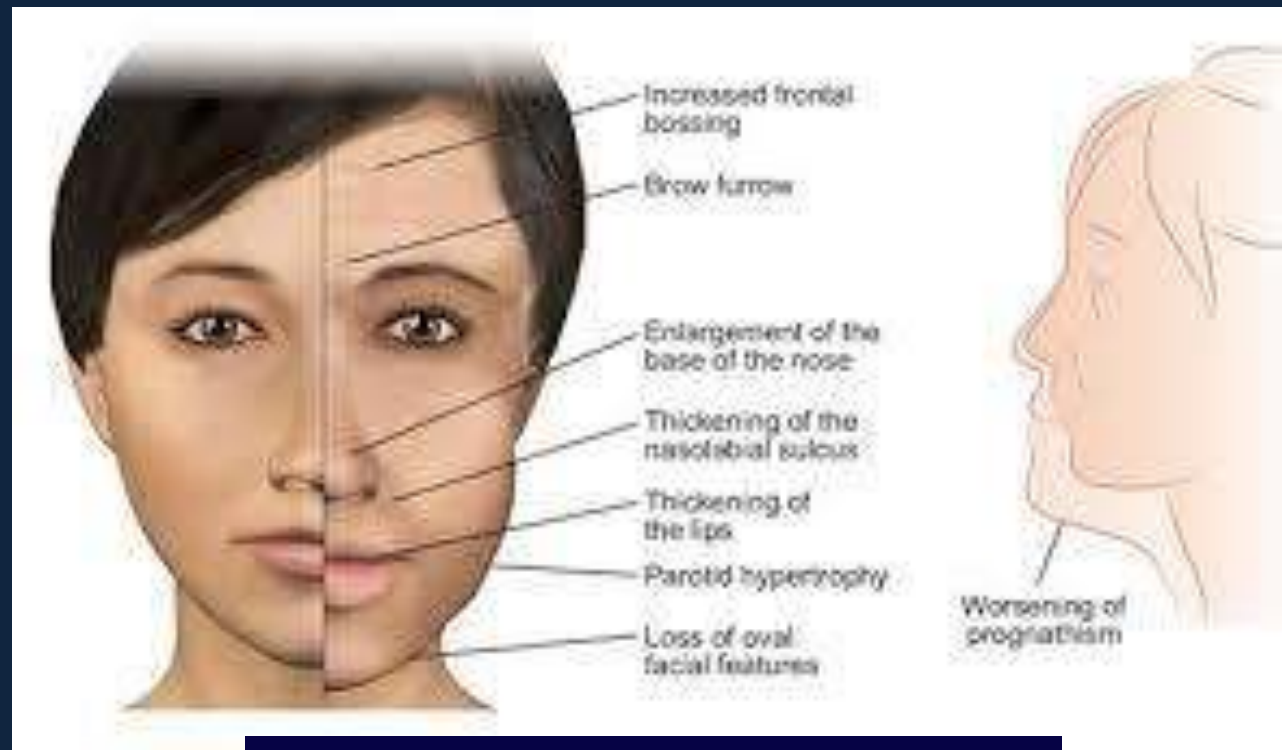
SP

Yudi Bijak Laksono 33268 (23 y , 22 y)
Reformat — REF COR POST C
33268
1004



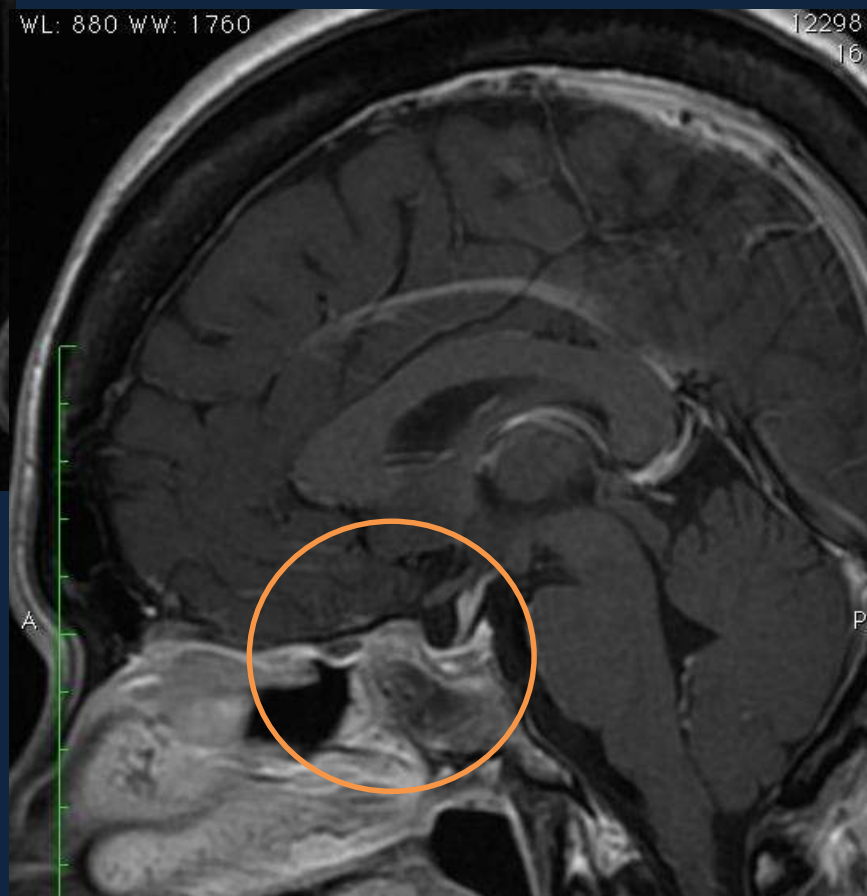
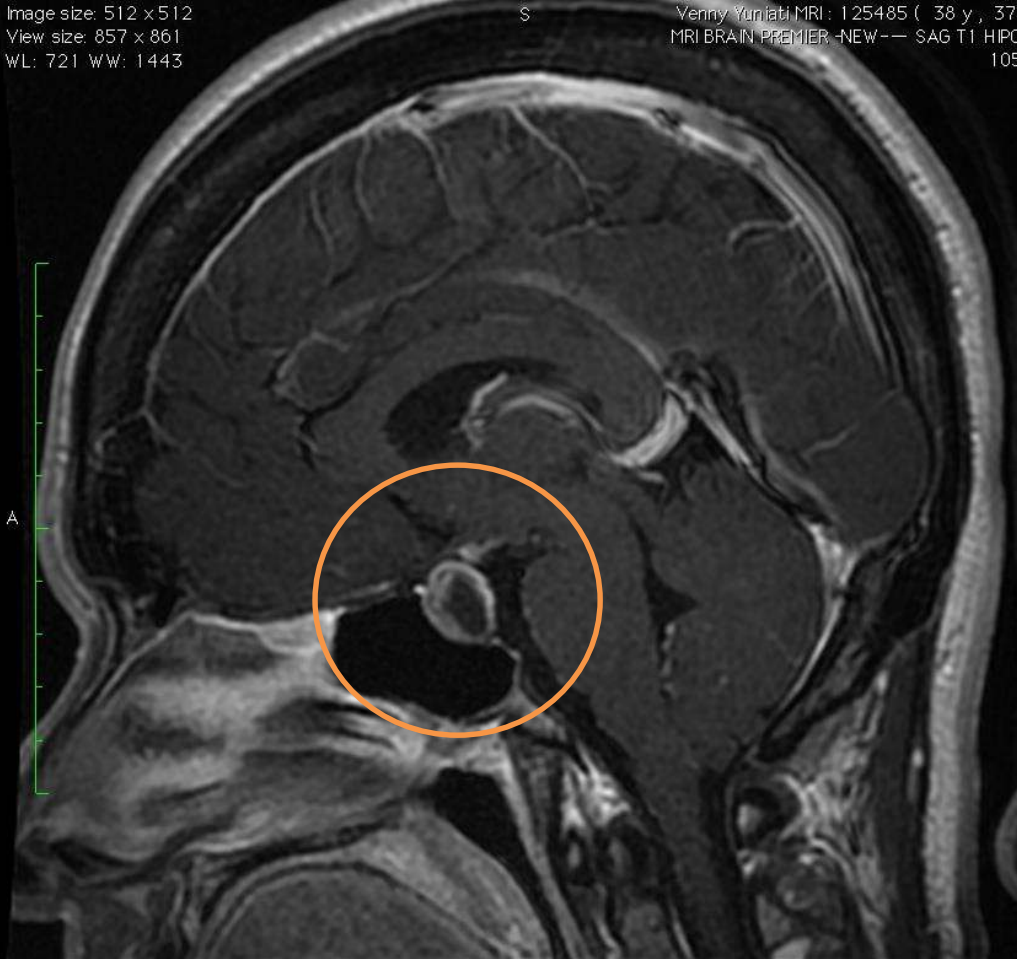
Pria, 23 tahun,
Gangguan Penglihatan (lapang pandang)
Gangguan ereksi
Peningkatan Hormon PROLAKTIN →
PROLACTINOMA

- Wanita, 34 tahun,
- gangguan menstruasi,
- tangan dan kaki makin membesar
- Peningkatan Hormon Pertumbuhan



ACROMEGALY

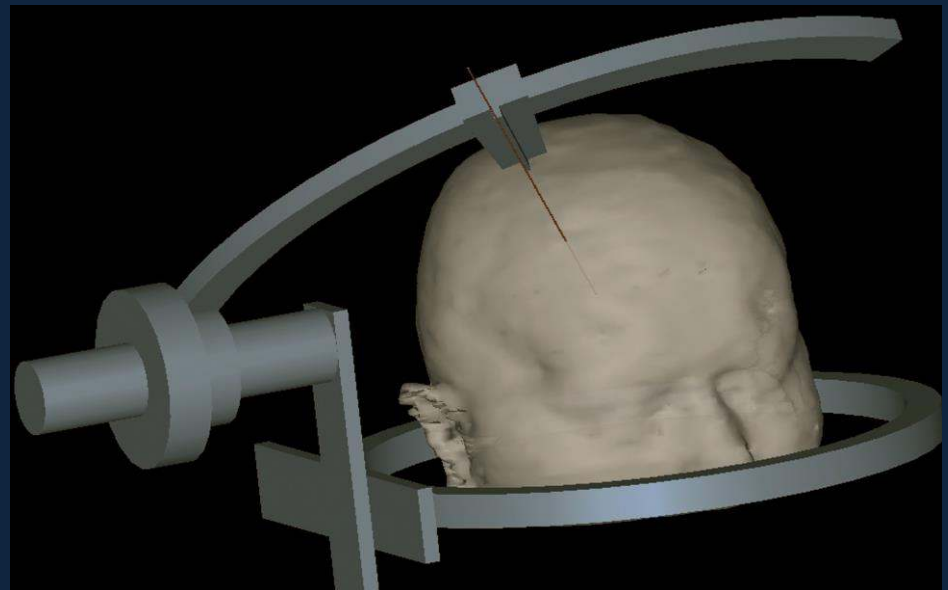
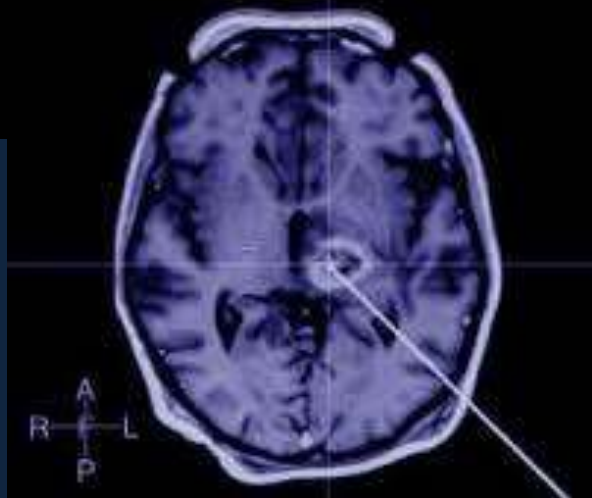
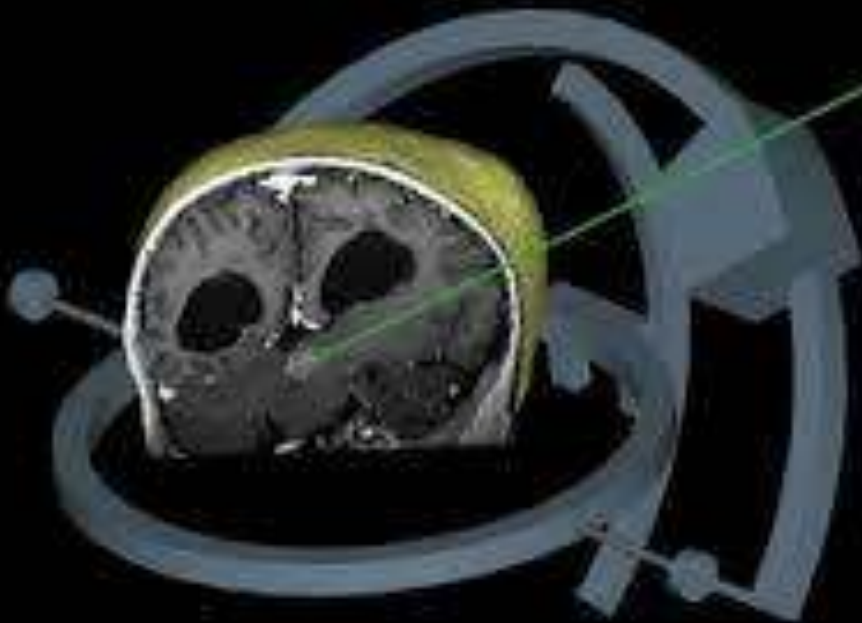




**Wanita, 34 tahun,
gangguan menstruasi,
tangan dan kaki makin membesar
(ACROMEGALY)
Peningkatan Hormon Pertumbuhan**

TUMOR OTAK DI LOKASI YANG DALAM

STEREOTACTIC

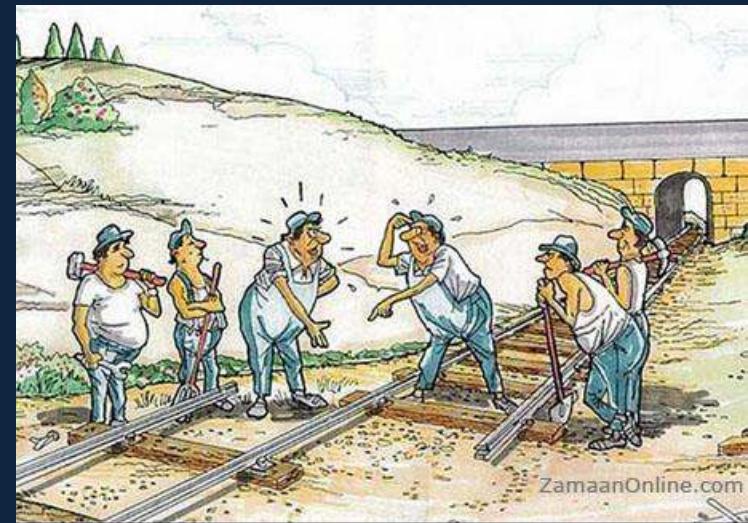


KEUNGGULAN TEHNIK MINIMAL INVASIVE

- Rasa sakit lebih sedikit
- Tidak ada / meminimalisir bekas luka pada wajah dan kulit kepala
 - Kosmetik
 - Scar minimal
- Masa pemulihan lebih cepat
 - **WAKTU DI RUMAH SAKIT LEBIH SINGKAT**

TIM DOKTER komprehensif:

- Neurosurgeon
- Neurologist
- Endokrinologist
- Neuro – radiologist
- Neuro Patologist
- Ophthalmologist
- Medical Rehabilitation
- Perawat terlatih khusus



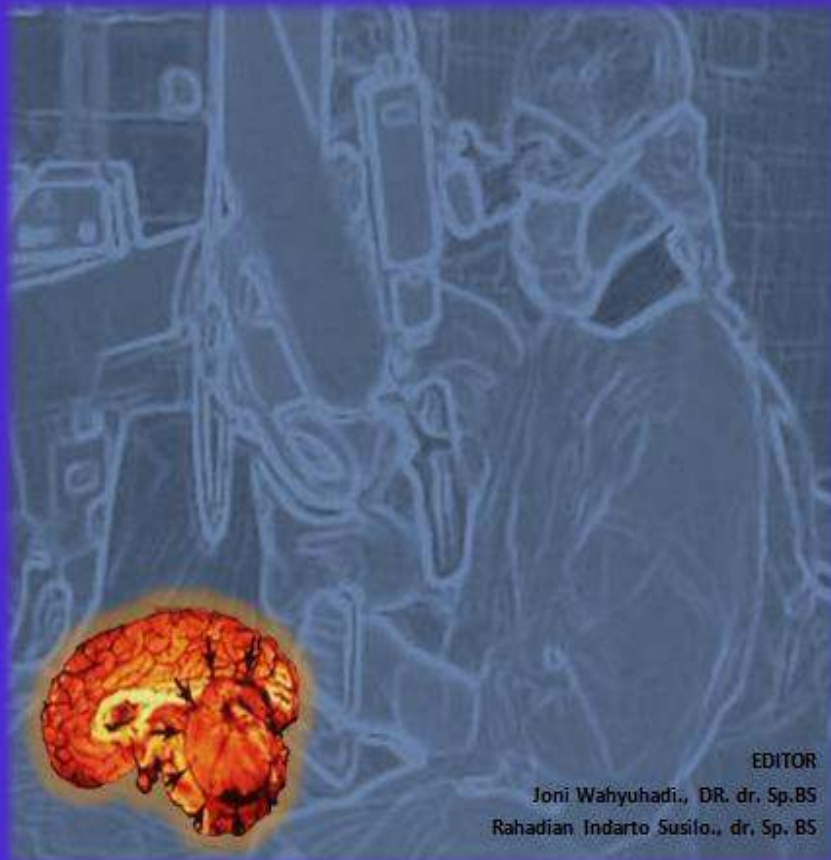
TERIMA KASIH

**KEEP YOUR
BRAIN SAFE**

BrainTumorNetwork
BrainTumorIndonesia



**PEDOMAN TATALAKSANA
TUMOR OTAK**
(Guideline for Management of Brain Tumors)



EDITOR

Joni Wahyuhadi., DR. dr. Sp.BS
Rahadian Indarto Susilo., dr. Sp. BS

DIVISI NEUROONKOLOGI

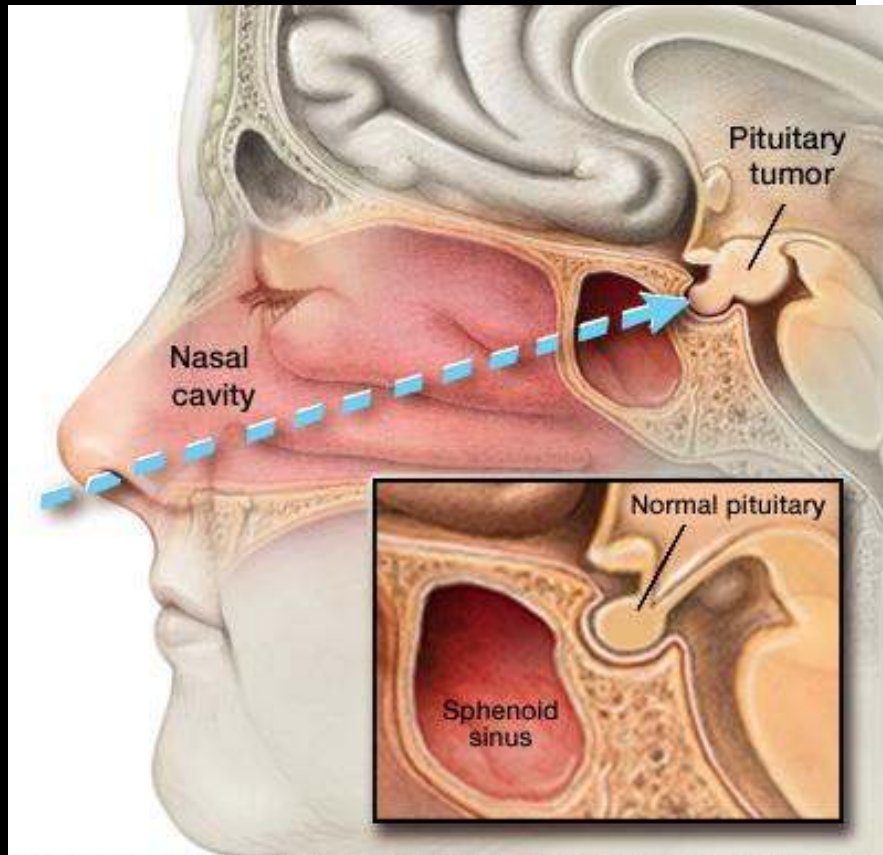
DEPT/SMF ILMU BEDAH SARAF
FAKULTAS KEDOKTERAN UNIVERSITAS AIRLANGGA
RSUD.Dr.SOETOMO SURABAYA
2010

Our Guideline

Microscope

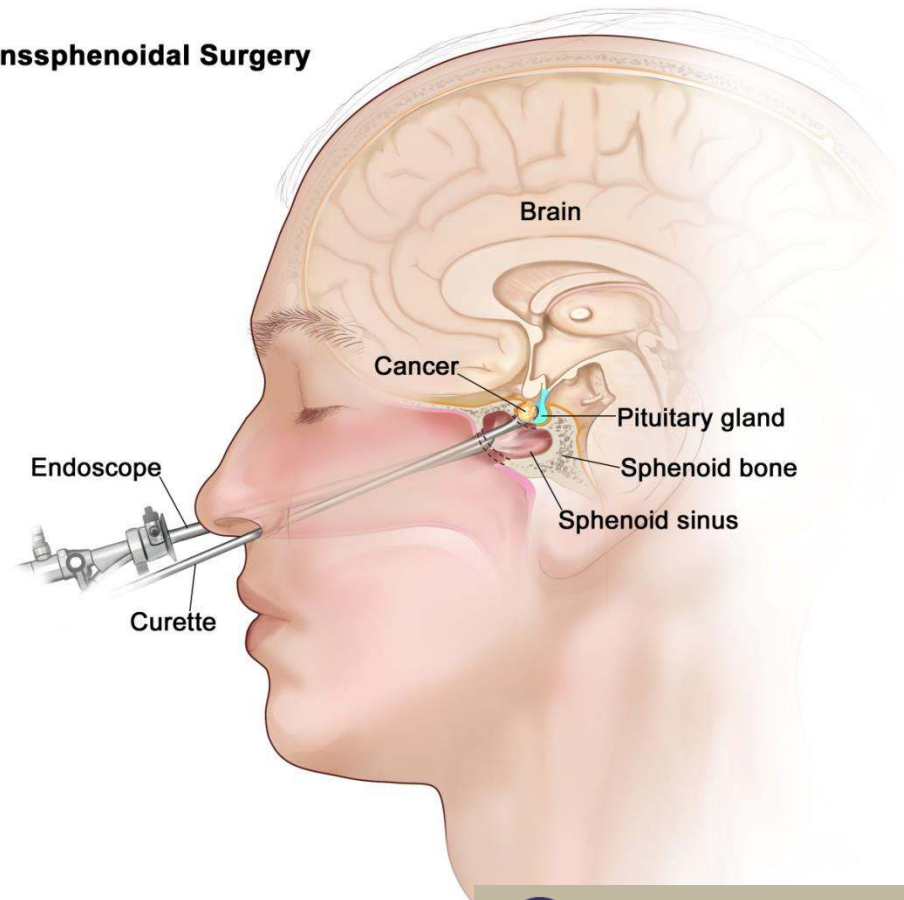


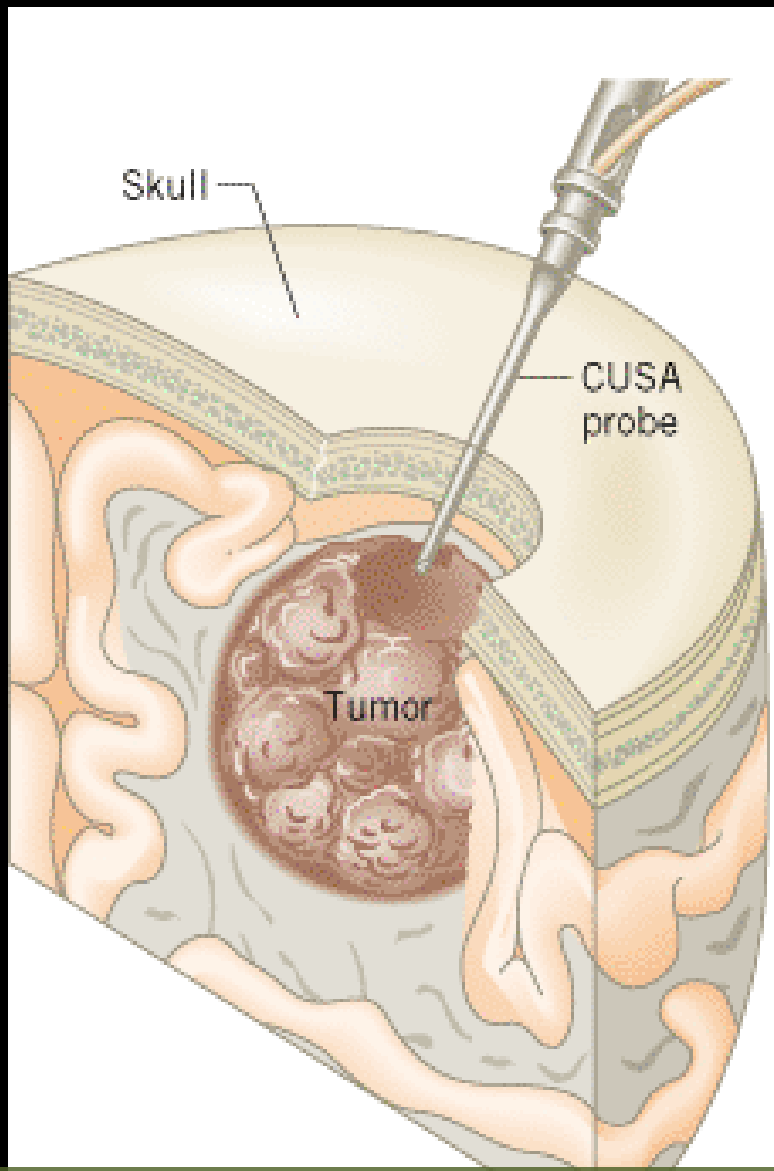
Endoscopy neurosurgery



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Transsphenoidal Surgery





cavitron ultrasonic surgical aspirator (CUSA)

Intraoperative monitoring

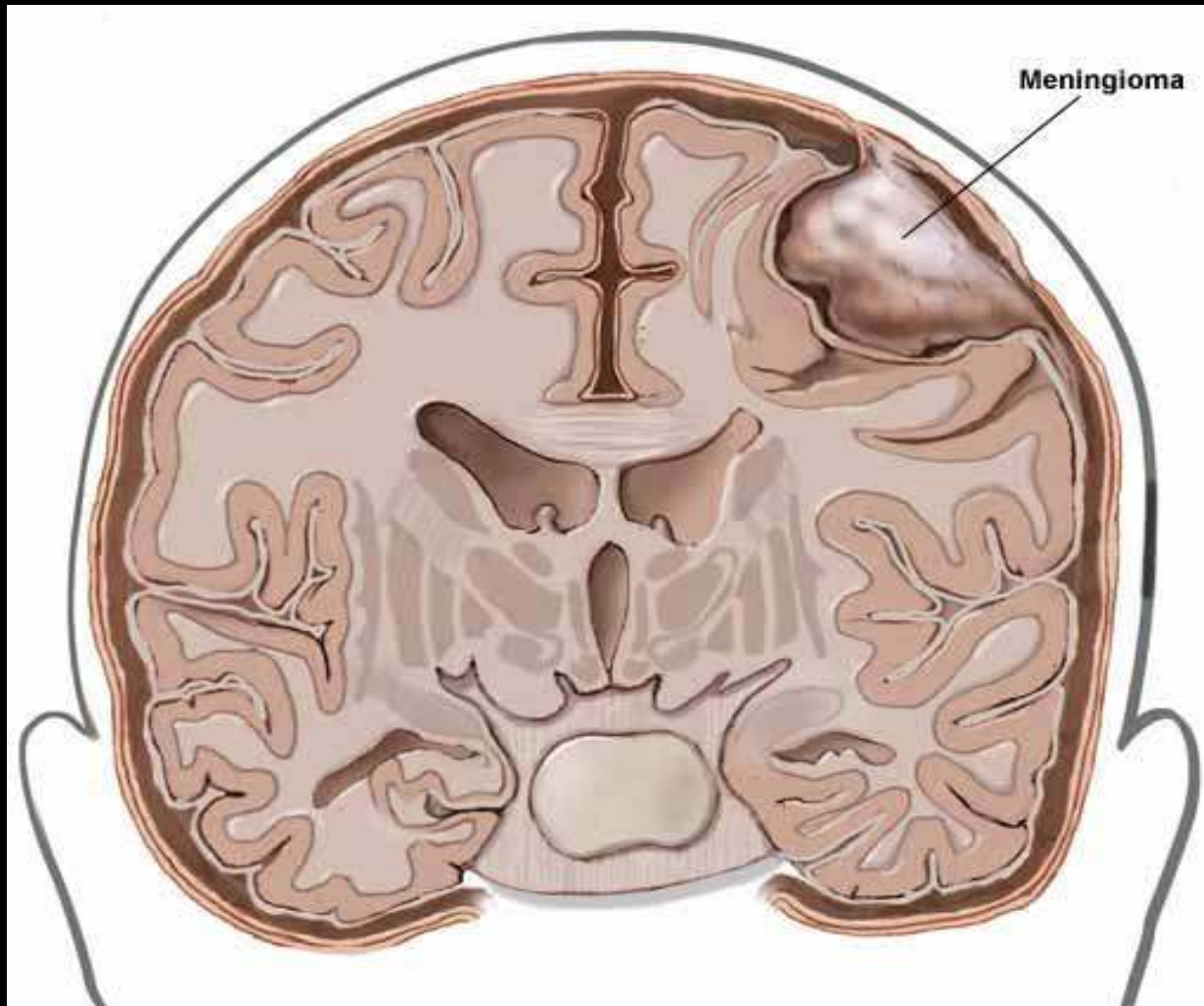
- ▣ Cranial nerve
- ▣ Brain Mapping
- ▣ Awake surgery

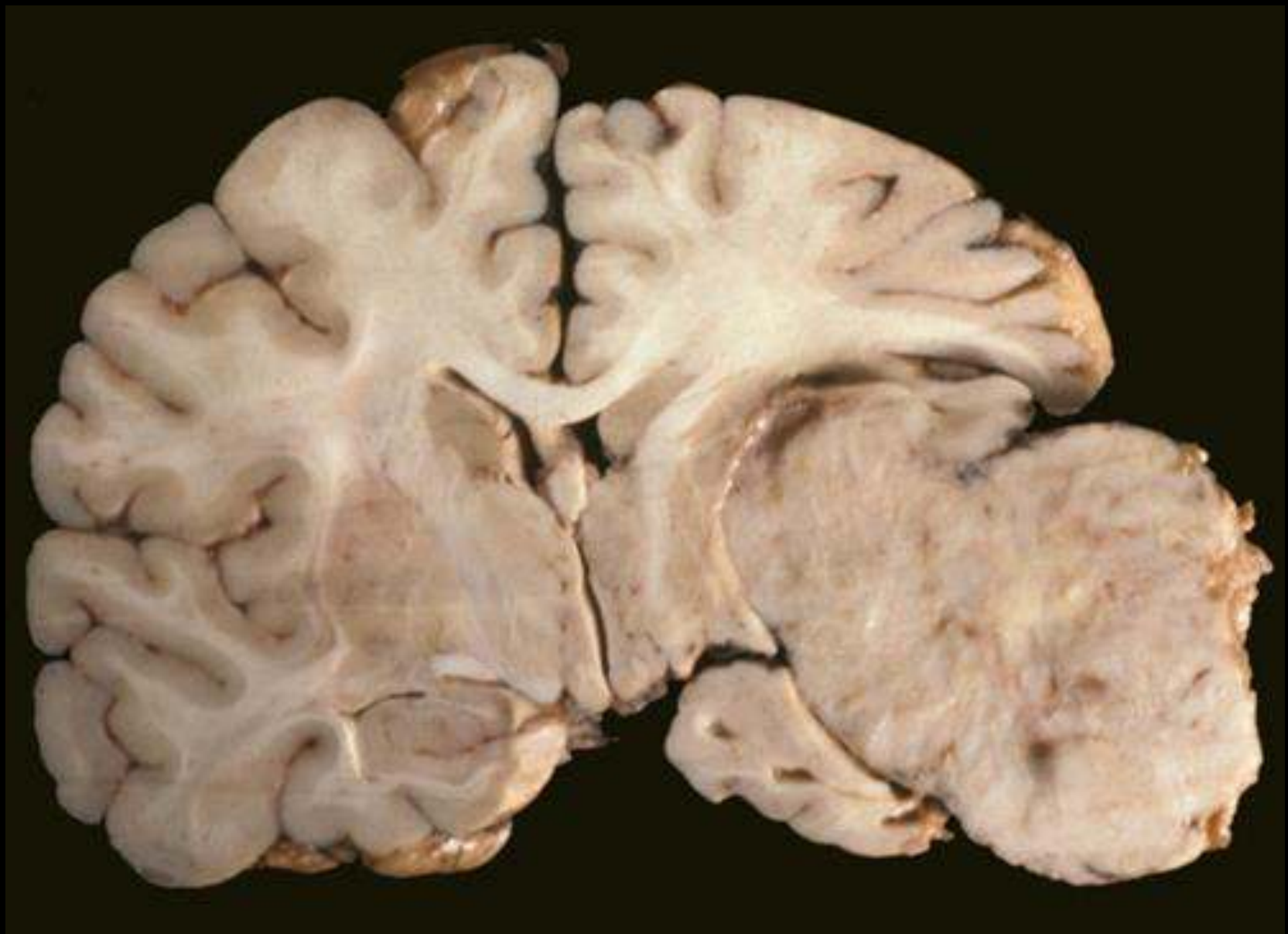


FOCUS

1. Meningioma
2. Astrocytoma
3. Metastase Brain tumor
4. Acoustic Schwanoma
5. Pituitary Adenoma

MENINGIOMA





Meningioma

- ▣ Origin : arachnoid cap cell → extra-axial tumor
- ▣ Slow growing
 - Incidental discovered meningioma → 32 % not grow over 3 yrs follow up
- ▣ Often produces hyperostosis of adjacent bone

Location

- ▣ Sphenoid wing (ridge) meningioma
- ▣ Parasagittal and Falk meningioma
- ▣ Olfactory groove meningioma
 - Tumor > 3 cm → The morbidity, mortality and difficulty in achieving total removal increase significantly
- ▣ Planum sphenoidale meningioma
- ▣ Tuberculum sellae meningioma
- ▣ Foramen magnum meningioma
- ▣ Convexity

SURGICAL INDICATION

- ▣ :Documented growth on serial imaging
- ▣ Symptoms referable to the lesion that no satisfactory controlled medically
- ▣ **GOAL → complete removal**
(tumor + dural attachment + abnormal bone)

IN: 6619
Se: 4
SN 587.75
In: 25+C

DI0V 25.0cm
SIND/L/



ANISA, N
I 39 400126

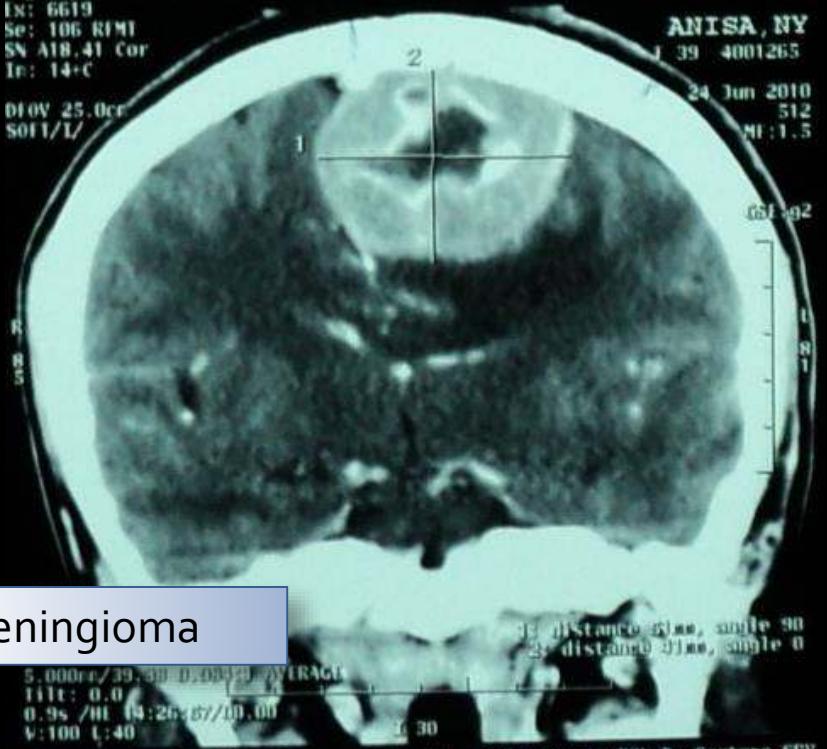
24 Jun 20
5
MI:1

GSE:



LightSpeed VCI SYSD199
IN: 6619
Se: 106 R1M1
SN A18.41 Cor
In: 14+C

DI0V 25.0cm
S011/L/



ANISA, NY
I 39 4001265

24 Jun 2010
512
MI:1.5

651.92

1: distance 31mm, angle 90
2: distance 31mm, angle 0

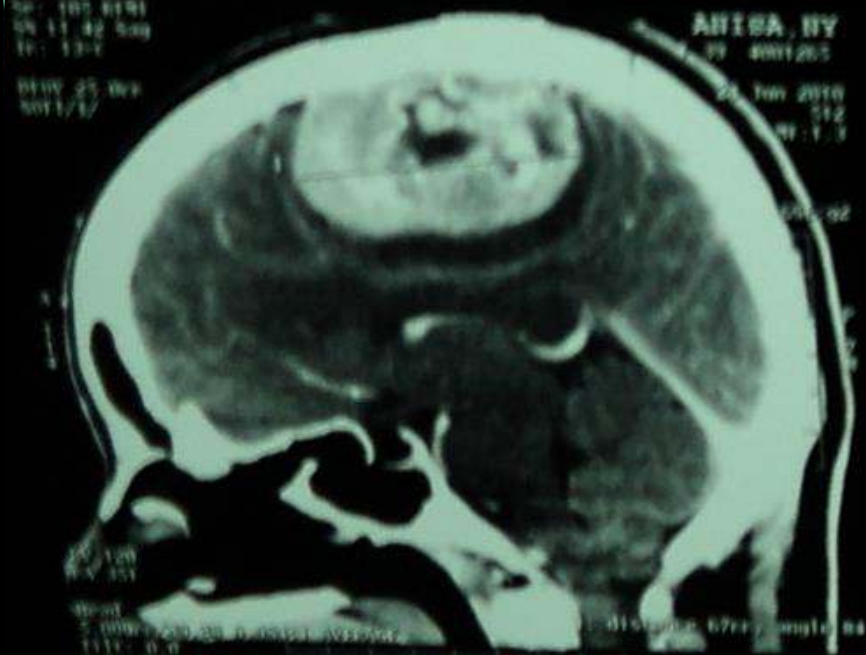
5.000mm/39.50 D:00000 AVERAGE
1111: 0.0
0.9s /HL 14:26:57/10.00
V:100 L:40

LightSpeed VCI SYSD199

S 144 Graha Arerta RSU Dr Soetoro SEV

LV 120
IN: 6619
Se: 106 R1M1
SN 11.42 Sag
In: 13+C

DI0V 25.0cm
S011/L/



ANISA, NY
I 39 4001265

24 Jun 2010
512
MI:1.5

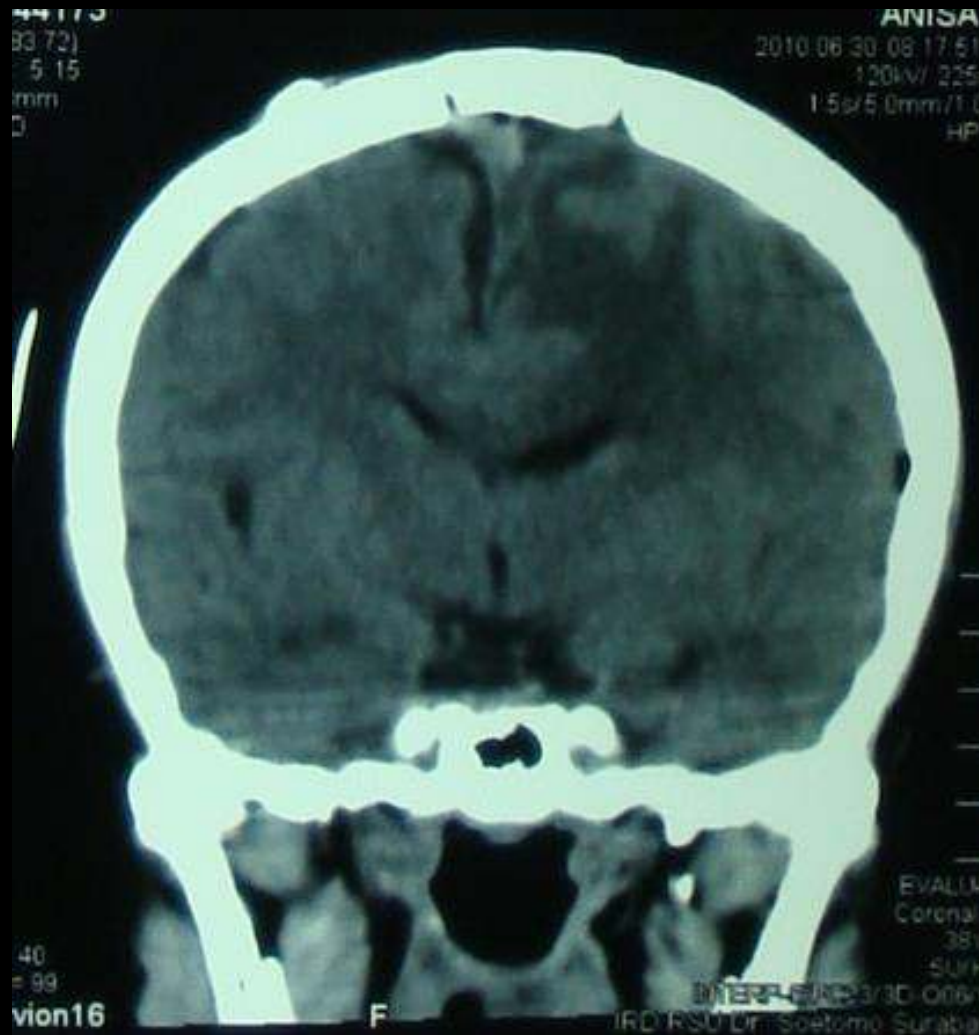
651.92

1: distance 67mm, angle 84

5.000mm/39.50 D:00000 AVERAGE
1111: 0.0
0.9s /HL 14:26:57/10.00
V:100 L:45

V:100 L:45

F 40 yo/ Parasagittal meningioma



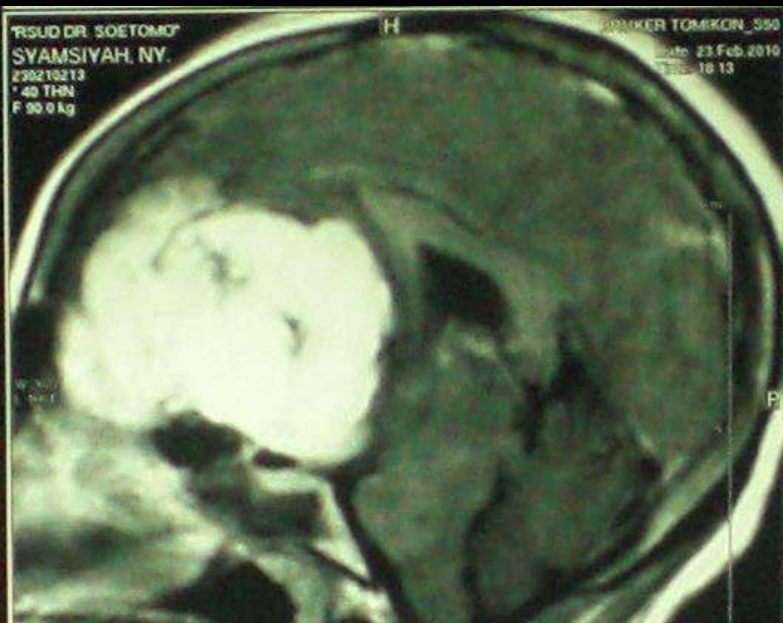
GROSS TOTAL REMOVAL

Olfactory Groove Meningioma

Tumor > 3 cm → The morbidity, mortality and difficulty in achieving total removal increase significantly

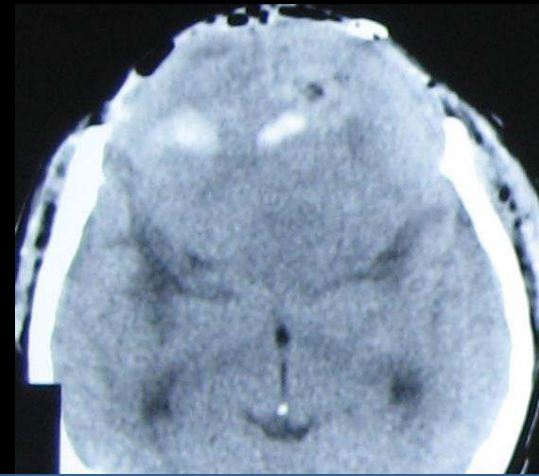


GROSS TOTAL REMOVAL



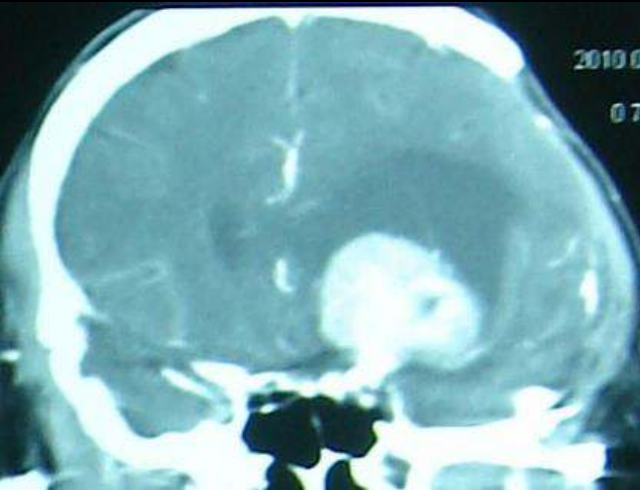
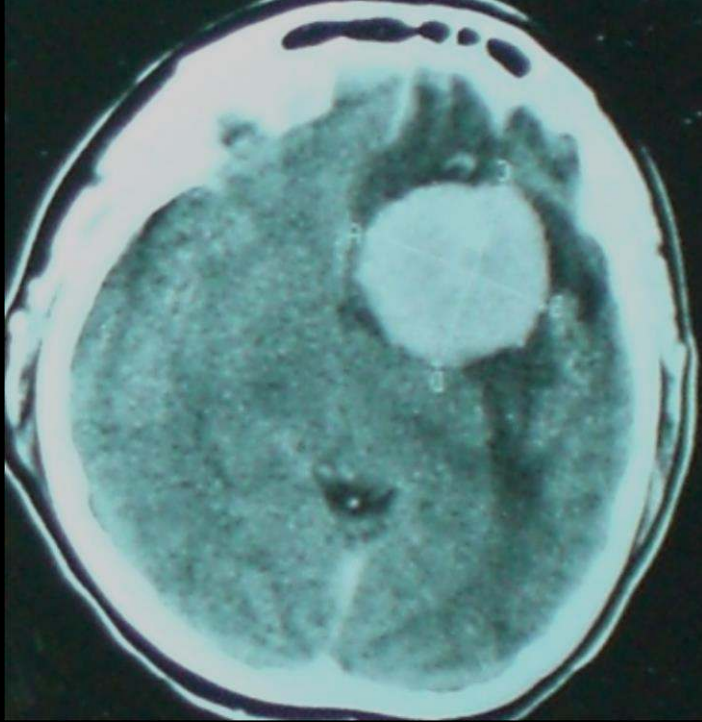
Olfactory Groove Meningioma meluas sampai ke posterior

Tumor > 3 cm → The morbidity, mortality and difficulty increase significantly



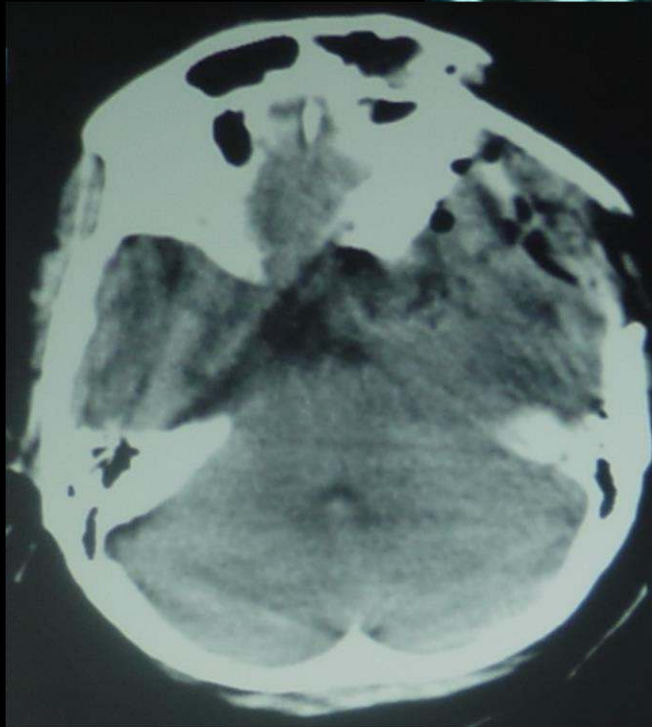
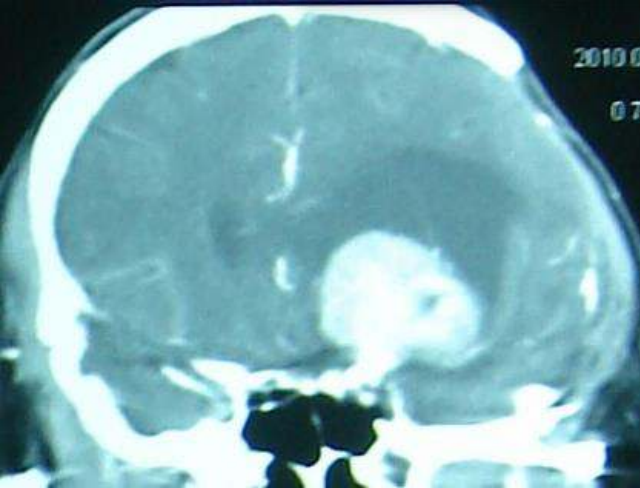
GCS menurun → Op. emergency at IRD → simple decompressi

Sphenoid medial / clinoidal meningioma

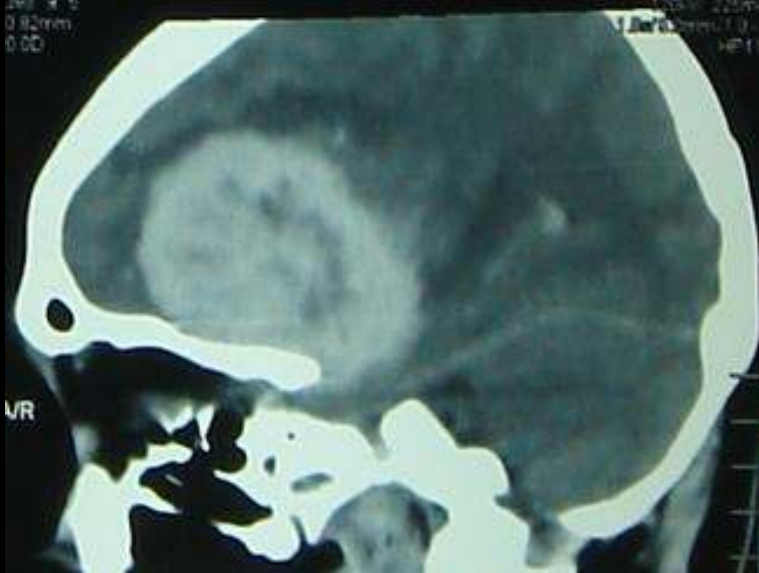


op I (in other institution) → only partial excision

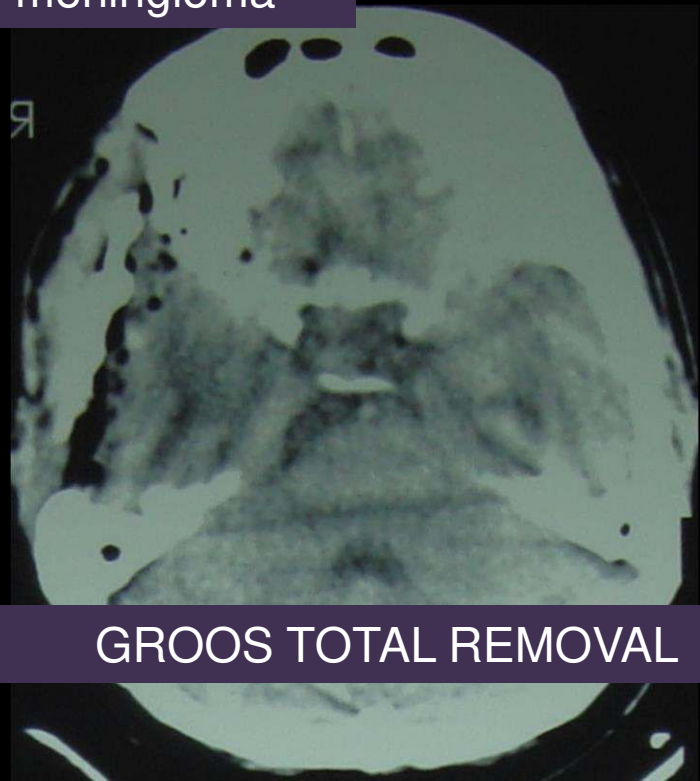
RESIDUAL Sphenoid medial / clinoidal meningioma



Re – operasi → Hasil op 2
→ gross total removal



F 44 yo/ Sphenoid medial meningioma

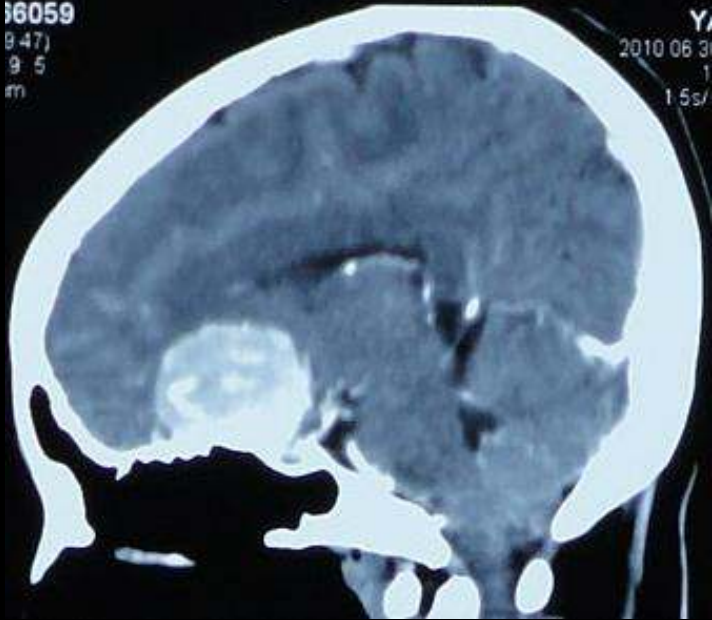


GROOS TOTAL REMOVAL

16059

8:47
9 5
m

Y)
2010 06 31
1
1.5s/



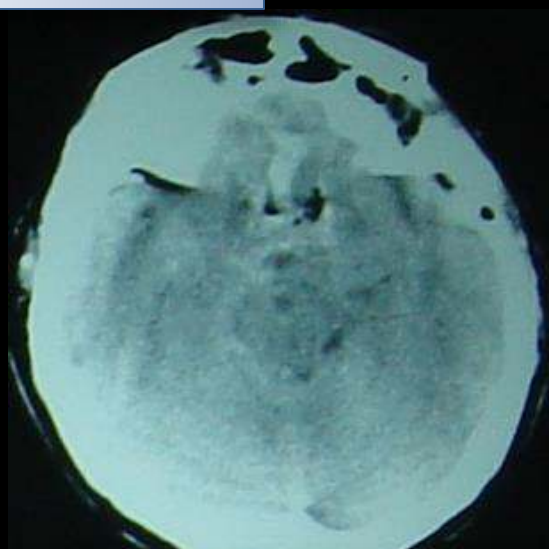
2010 06 30 08
120kV
1.5s/5.0m



TUBERCULUM SELLAE MENINGIOMA

(178.85)
#1 7.6
76mm
00

2010 06 30 08 06 4
120kV/ 22
1.5s/5.0mm/1
H



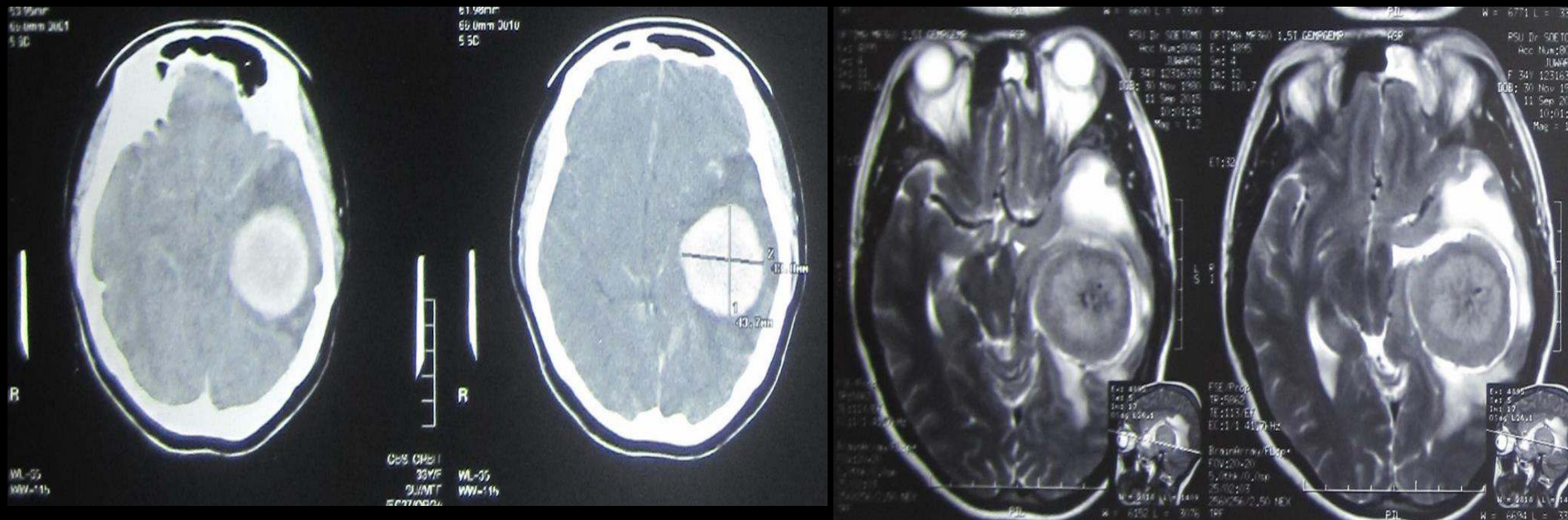
GROSS TOTAL REMOVAL

1:40
12 99

Post Op

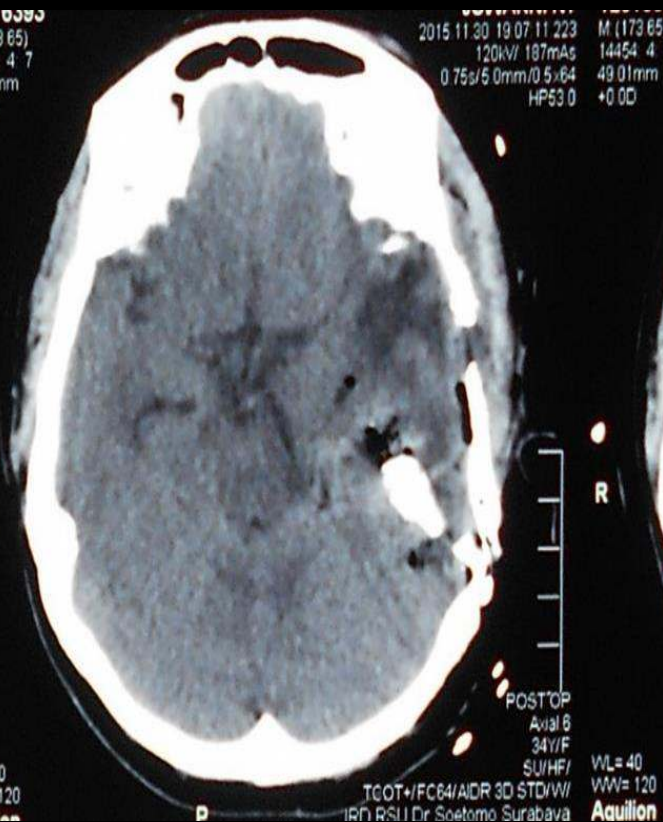
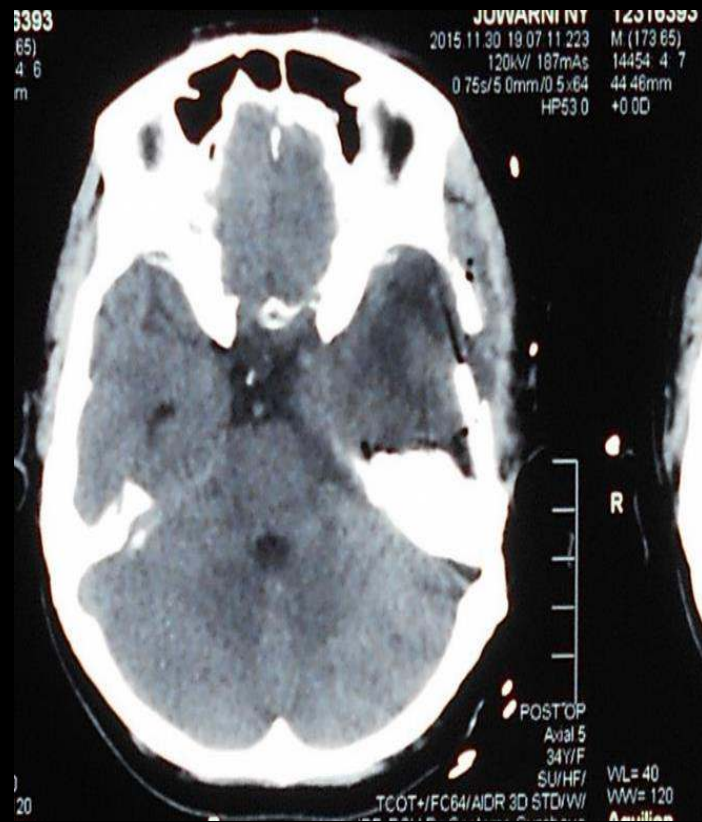


Meningioma Petrosal (S)

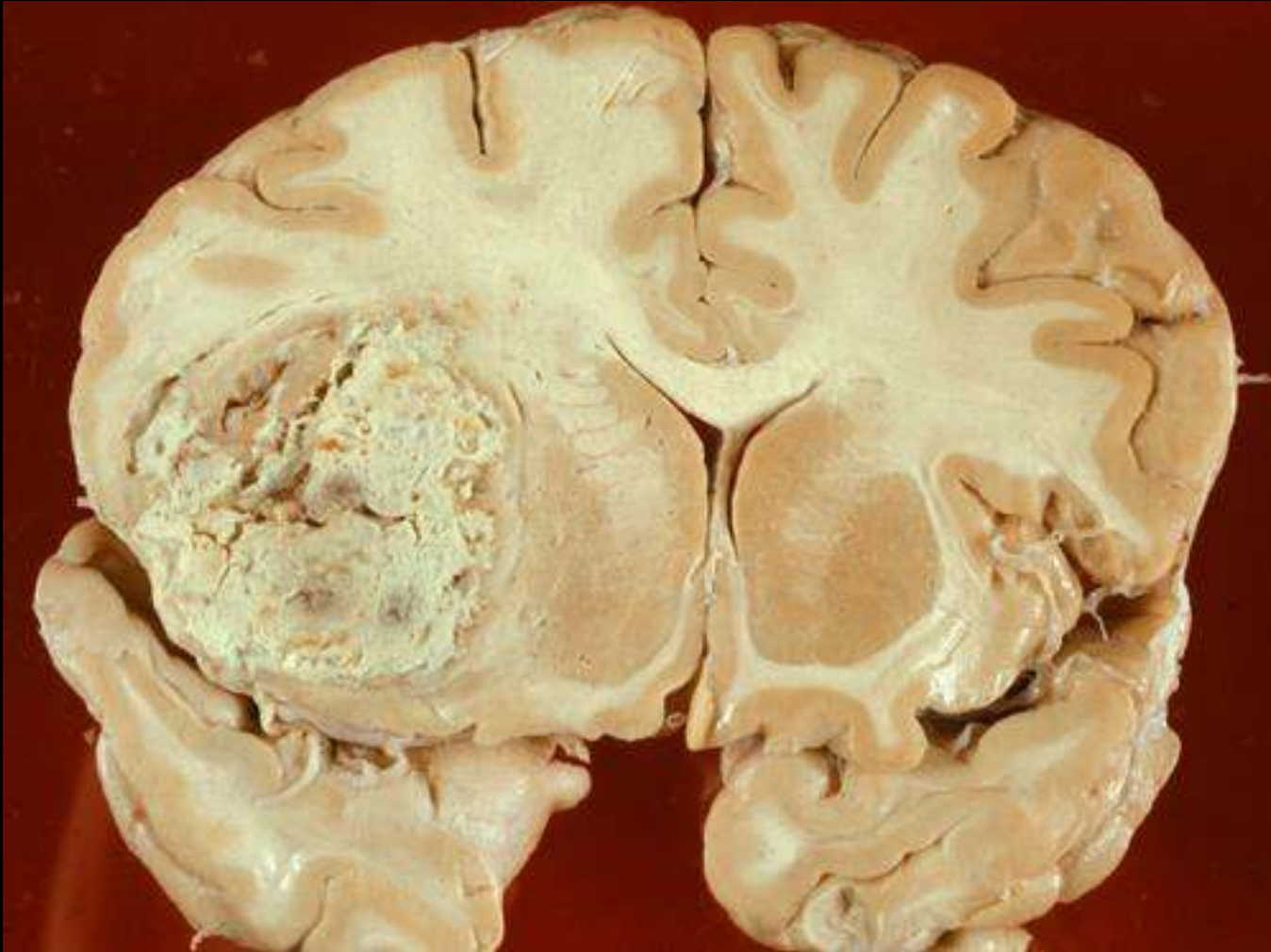


Perempuan 34 th/ pandangan mata kiri kabur sejak 6 bulan, nyeri kepala + / GCS 15 VOD 1/60, VOS 1/300, ODS early papil edema

Post Op



ASTROCYTOMA



Epidemiology

- ▣ **Most primary brain tumours are GLIOMA**
 - astrocytic
 - oligodendroglial
 - mixed astrocytic & oligodendroglial
 - ependymal neoplasms

- ▣ **Astrositoma**
 - The most invasive
 - Most commonly occurring brain tumors in all age

- ▣ **Canada in 2008:**
 - approximately 2550 cases (1450 men & 1100 women) diagnosed with primary brain tumours
 - 1,740 men and women will die of the same.

WHO Classification

- ▣ Grade I (Pilocystic Astrocytoma)
 - Most frequent brain tumors in children
- ▣ Grade II (low grade Astrocytoma)
 - 25% of glioma & infiltrative
- ▣ Grade III (anaplastic Astrocytoma)
 - Highly malignant , progress to GBM
- ▣ Grade IV (glioblastoma multiforme)
 - Highly malignant, poor prognosis
 - rapid spread to other region of the brain



**Low
grade**



**High
grade**

Clinical features

Clinical Features

- Low mitotic index
- Diffuse invasion
- High rate of transformation

**Low grade
Astrocytoma**

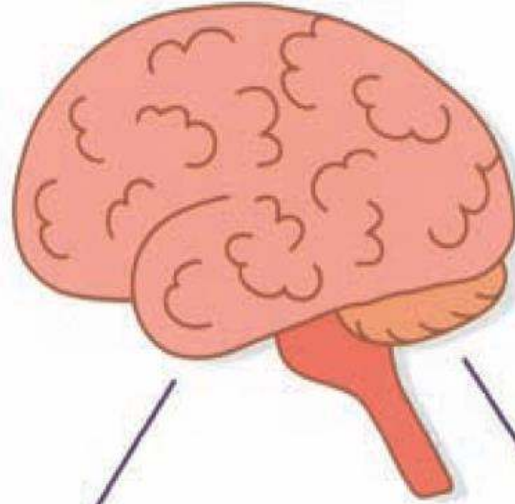
5-10 years

**Primary
Glioblastoma**

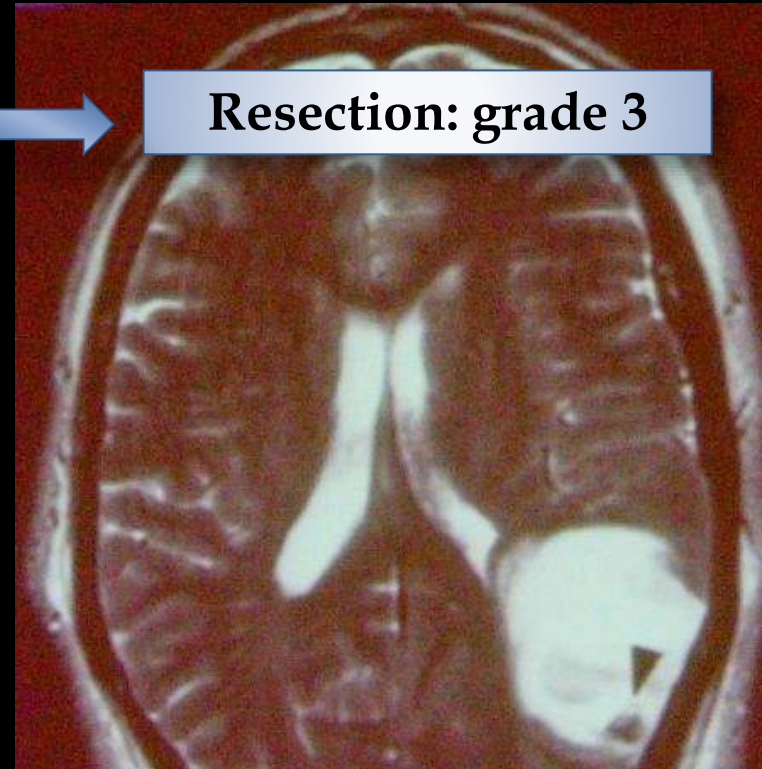
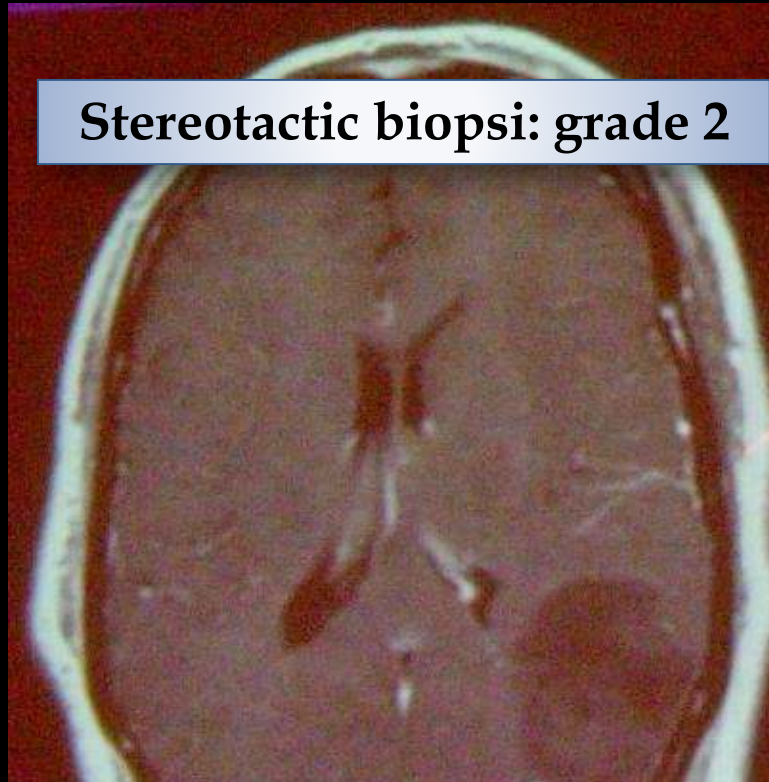
**Secondary
Glioblastoma**

Clinical Features

- Rapid proliferation
- Diffuse invasion
- Angiogenesis
- Cellular necrosis



Making an accurate diagnosis



81 patients sent for resection after biopsy →

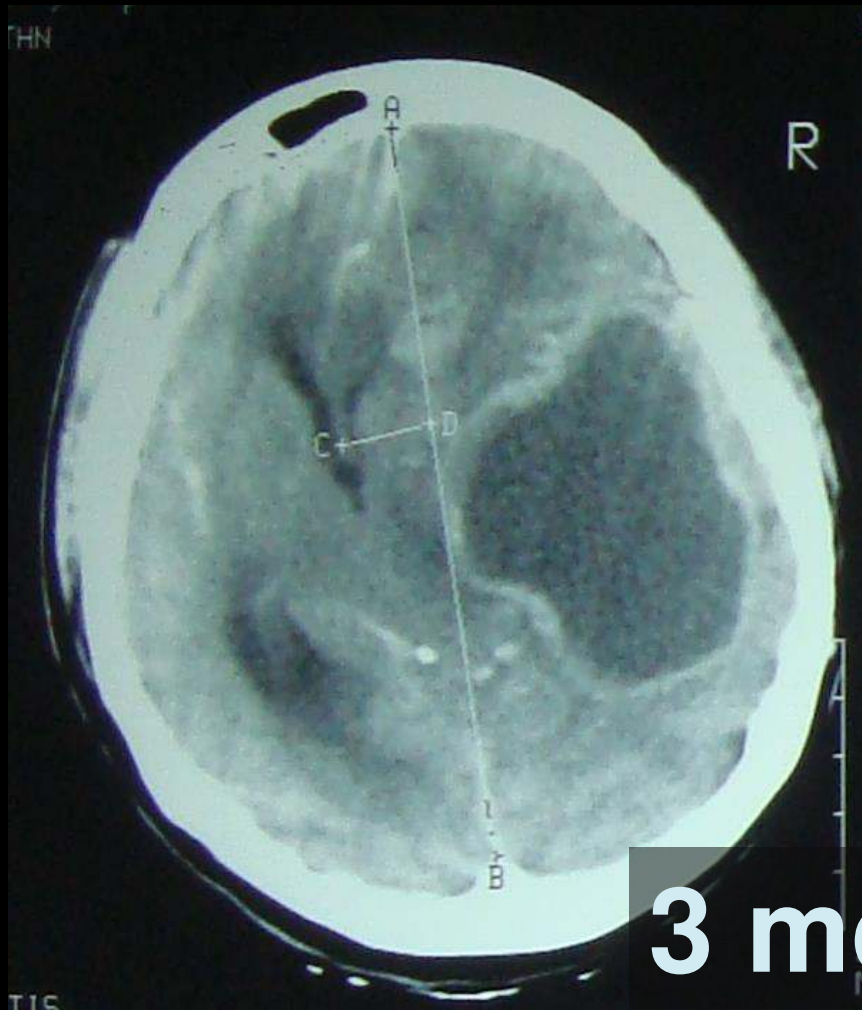
- 38% → change in pathology;
- 28% → this made a major difference in treatment

Major Prognostic Factors

- Age
- Karnofsky Performance Score
- Histological type
 - (oligodendroglial, mixed or astrocytic)
- Tumour grade
 - (WHO grade III or IV)

OTHER Prognostic factors :

- The extent of resection
- The location and volume of the tumour.



3 months

F/ 36 yo

Treatment

MULTIDISCIPLINARY

→ OPTIONS :

- No treatment → observation
- Surgery
- Radiation
- Chemotherapy

Tx High grade glioma

- ▣ **Surgery is the initial recommended approach** in suspected high grade gliomas,
- ▣ **Goal of surgery**
 - debulking,
 - clinical improvement
 - pathologic diagnosis
 - **SET UP OTHER THERAPIES**

SURGERY :
RESECTION vs BIOPSY

Surgery

- ▣ Surgical resection as the main treatment for
 - alleviation of symptoms and
 - prolongation of life
- ▣ Most studies report a survival advantage for resection over biopsy
 - Alberta Cancer Board, Clinical Practical Guideline, Management of Glioblastoma Multiforme, 2009
- ▣ **Recommendation:**
gross total resection, when safe,
in high grade gliomas
(WHO grade III and IV).

Adjuvant Tx

- ▣ **Adjuvant chemo-radiation tx:**
the standard of care following surgery.
- ▣ Whenever possible, surgery will be
 - followed by adjuvant **combined Temozolomide-based chemotherapy and radiation** → “Concurrent Treatment”
 - followed by adjuvant **Temozolomide** for a **total of 6 to 12 cycles.**

“Concurrent Treatment”

- ▣ low dose of chemotherapy (temozolomide) daily during radiation therapy
- ▣ With this approach survival at 2 years improved from 10% to 26%.
- ▣ Determination of MGMT may assist in determination of prognosis

“concurrent treatment” followed by monthly temozolomide cycles (six monthly cycles of temozolomide administered 5 days every 28 days) is the standard of care for patients with GBM.

Local delivery

- ▣ **Deliver therapy directly to the tumor site**
- ▣ **increase efficacy**
- ▣ **Reduce side effects of systemic therapy**
- ▣ **Use less of the therapeutic drug**
- ▣ **Delivery drug continuously, and increase patient compliance**

Elderly Px

- ▣ Elderly patients (≥ 60 years old) with a poor performance status (KPS ≤ 70) → adjuvant radiation tx alone, with or without tissue diagnosis
- ▣ Radiation Tx → abbreviated to 40 Gy in 15 fractions

Low Grade Glioma

- ▣ Treatment → controversial
- ▣ requires a multidisciplinary approach which can include surgery, radiation therapy, and chemotherapy.
- ▣ Survival & response rates are related to
 - Tumour type
 - ▣ Oligodendrogliomas or oligo- dominant mixed histology has better survival
 - Size (smaller size associated with better survival)
 - Age of the patient (younger is better),
 - Mental status

Tx Low Grade Glioma

- ▣ In patients with low grade glioma and controlled epilepsy as the single symptom, surgery may be deferred until clinical or radiological progression
- ▣ **maximal resection (when safe) if →**
 - In patients with increased intracranial pressure,
 - neurological deficits,
 - uncontrollable seizures,
 - clinical or radiological progression,

Radiotherapy in Low Grade Glioma

- ▣ **Postsurgical radiation therapy**
 - may be deferred until clinical or radiological progression
- ▣ When Radiation therapy is indicated, the dose should be between **45 and 54 Gy**

Chemotherapy in Low Grade Glioma

- ▣ Chemotherapy **should not be added** to radiation therapy
 - the combination shows no benefit in comparison to RT alone
 - increases the toxicity.
- ▣ Chemotherapy such as temozolomide may be offered in patients with **progressive tumours that harbour combined 1p/19q loss of heterozygosity.**

Difuse Astrocytoma (WHO grade II)

F/ 39 y.o
GCS 15
Headache



The “high risk“ low grade glioma

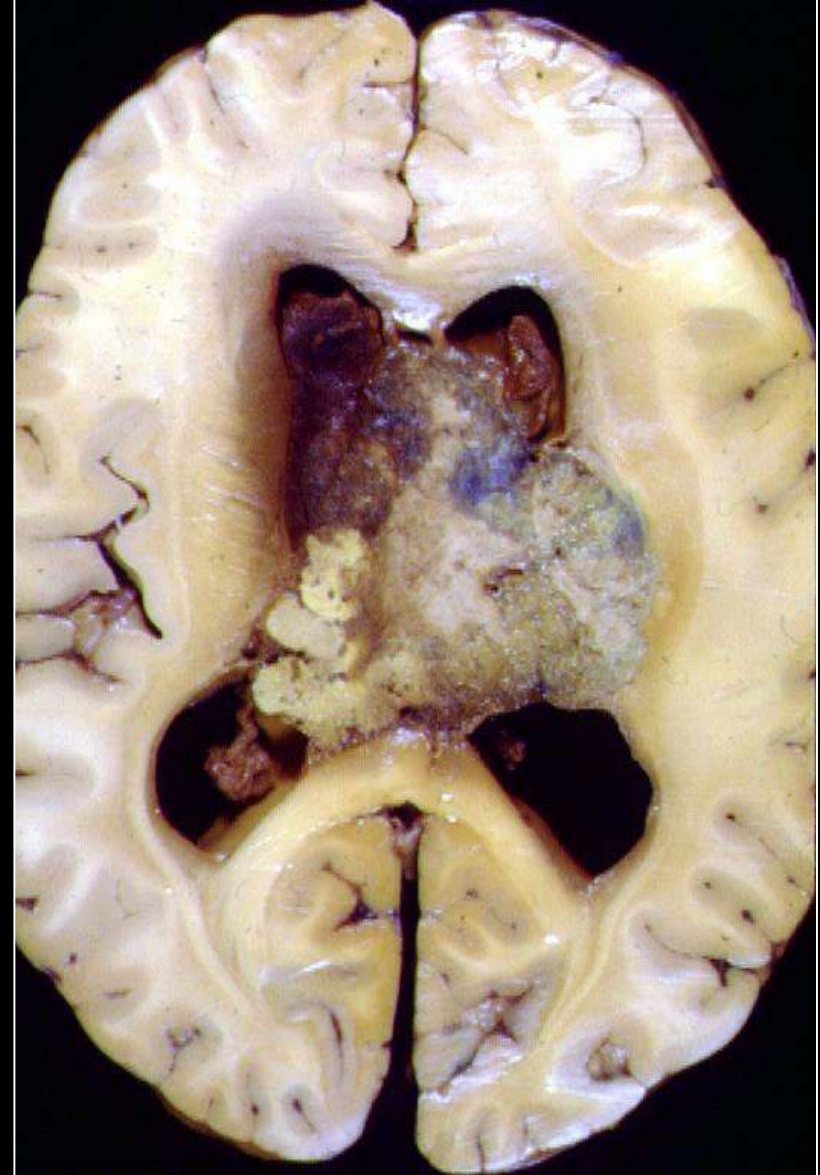
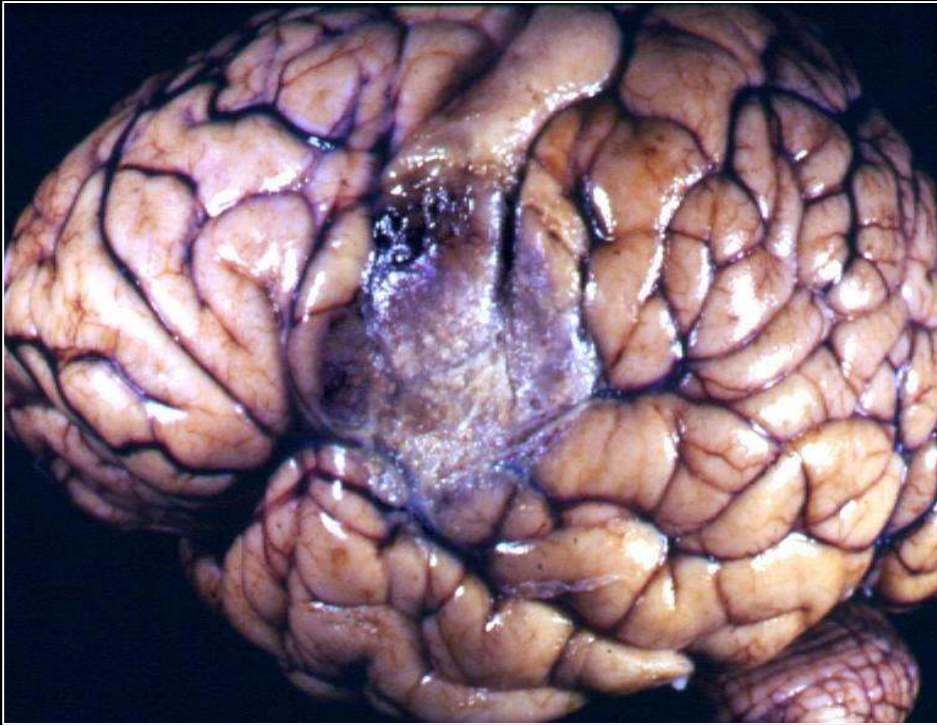
- ▣ Meet at least three of the following criteria :
 - Age ≥ 40 years,
 - Largest preoperative tumor diameter ≥ 6 cm
 - Tumor crossing midline
 - Tumor of astrocytoma histology
 - Preoperative neurologic deficits (Neurologic Function Score >1)
- ▣ Tx \rightarrow temozolomide concurrently with RT followed by 12 cycles of 4-weekly chemotherapy.
 - There is currently an ongoing phase II clinical trial

New Hope

- **Anti-Angiogenic Agents**
- **Gene Therapy**
- **Immunotherapy**
- **Molecular Pathway Inhibitors**

Metastatic Brain Tumor

Metastatic Tumors



25-30% of all intracranial tumours

Lung, breast commonest primary sites

Surgical Removal ??

- ▣ **Not all px** with brain metastases are candidates for resection
- ▣ Consider :
 - The lesion → Number, Location , Size
 - Clinical features of the px
 - Histology of primary tumor

Patient selection

▣ STATUS OF SYSTEMIC DISEASE

- Control of primary cancer → expected survival > 4 months
- General medical condition → able to withstand surgery / anesthesia
- Neurologic status → KPS > 70

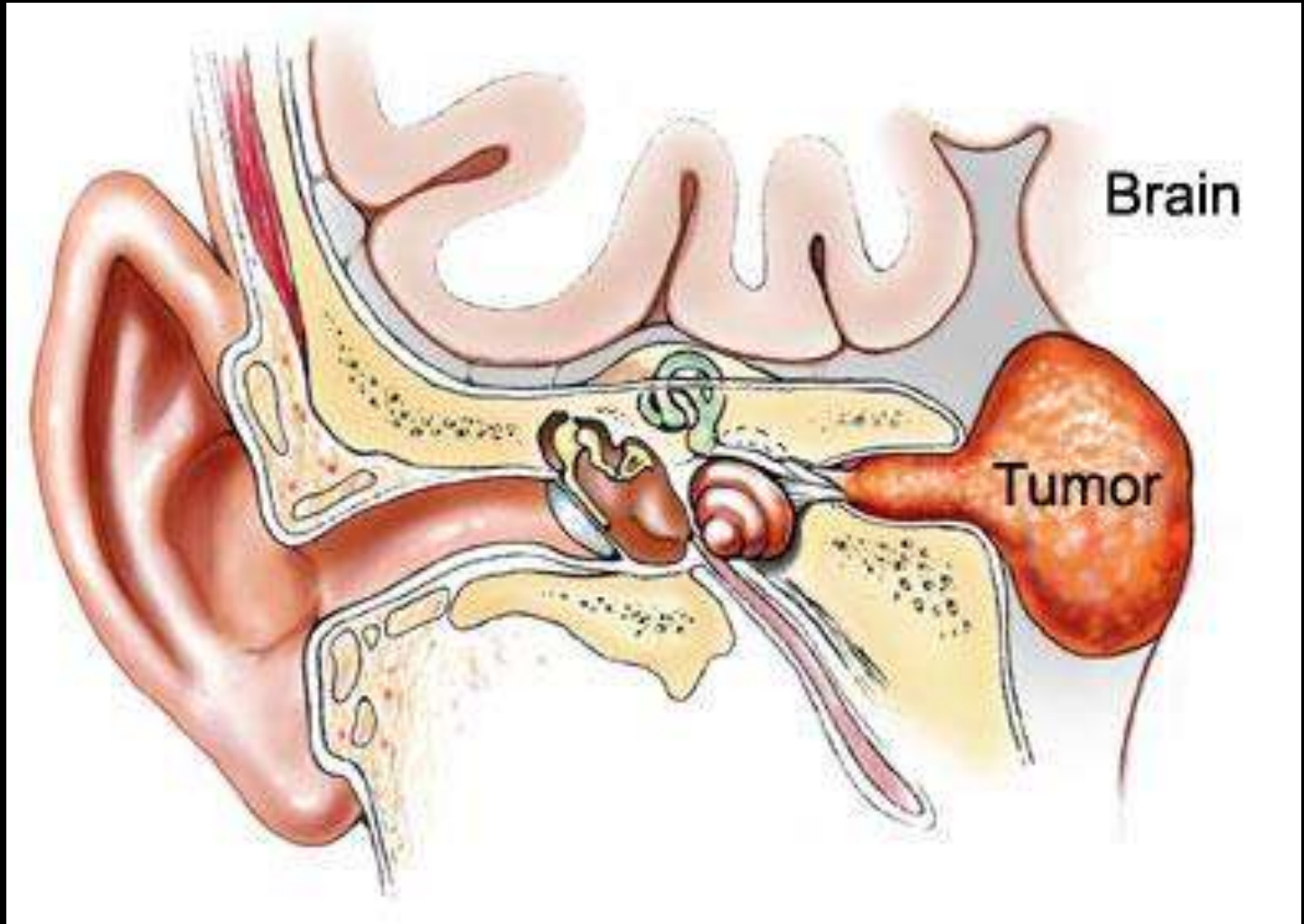
▣ RESECTABILITY

- Accessibility → not brain stem, basal ganglia, thalamus
- Numbers of lesions → ≤ 3

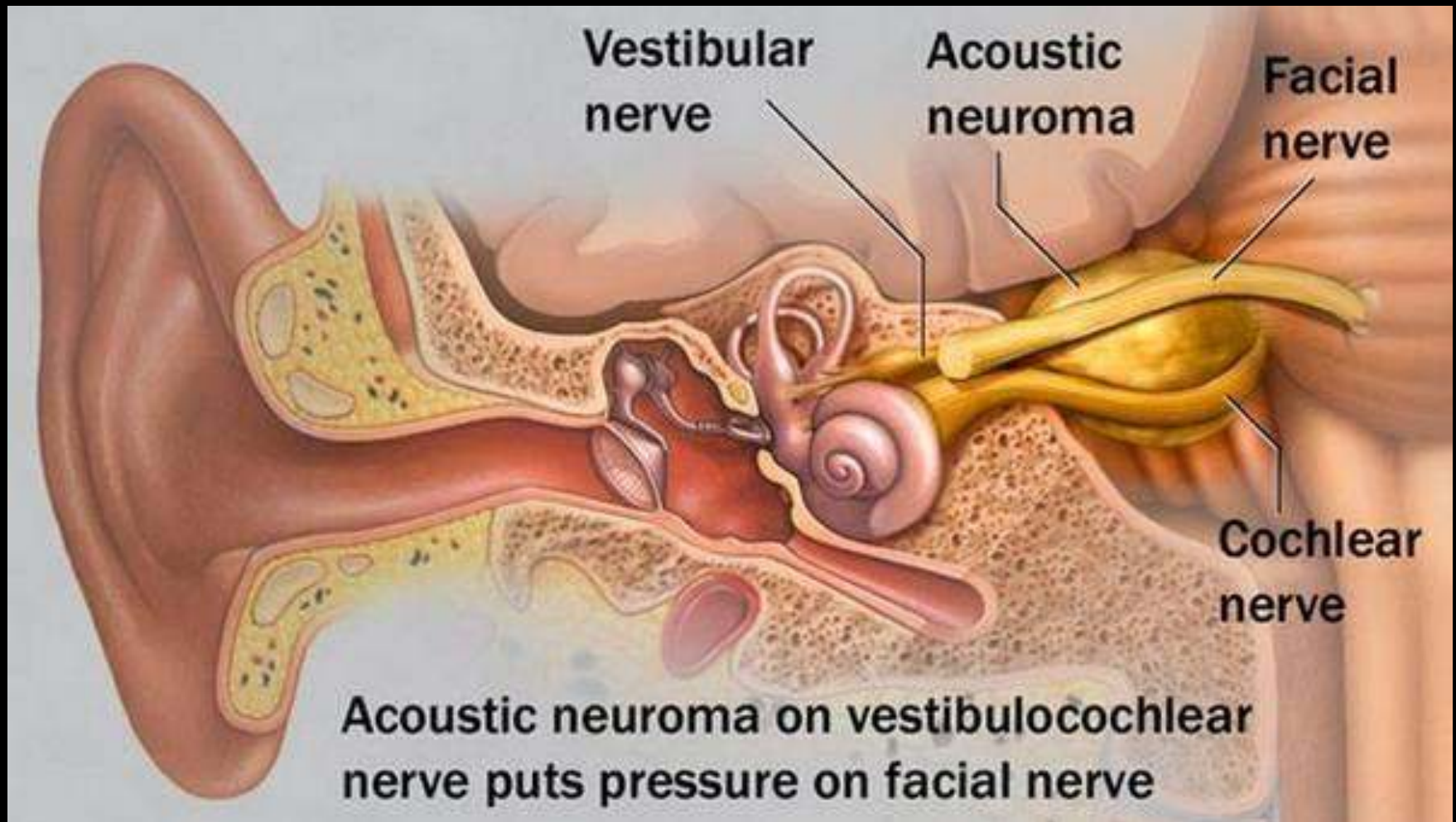
▣ RADIOSENSITIVITY

ACOUSTIC SCHWANNOMA

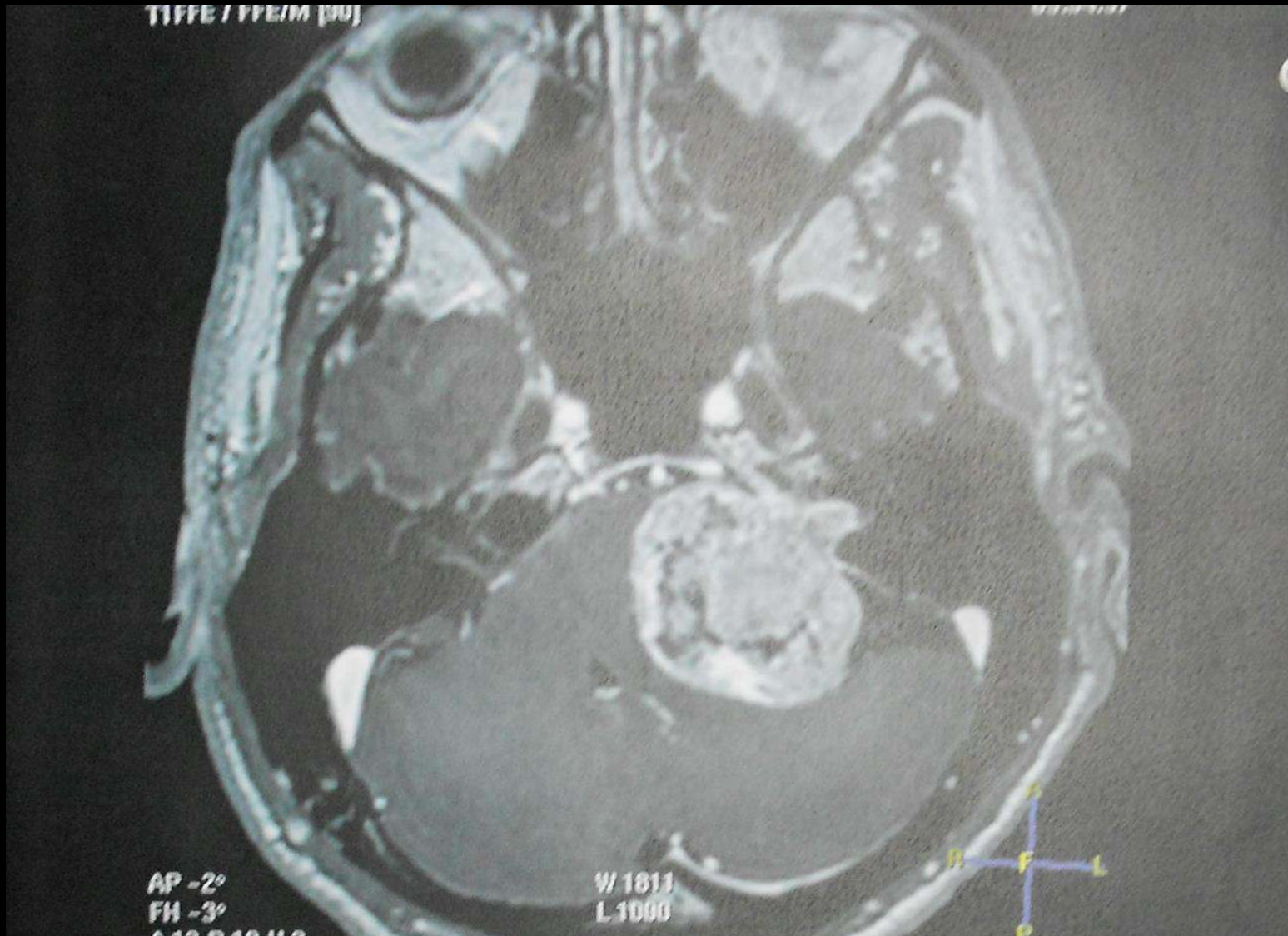
Acoustic schwannoma



Vestibular Schwannoma = Acoustic neuroma



Vestibular Schwannoma = Acoustic neurinoma



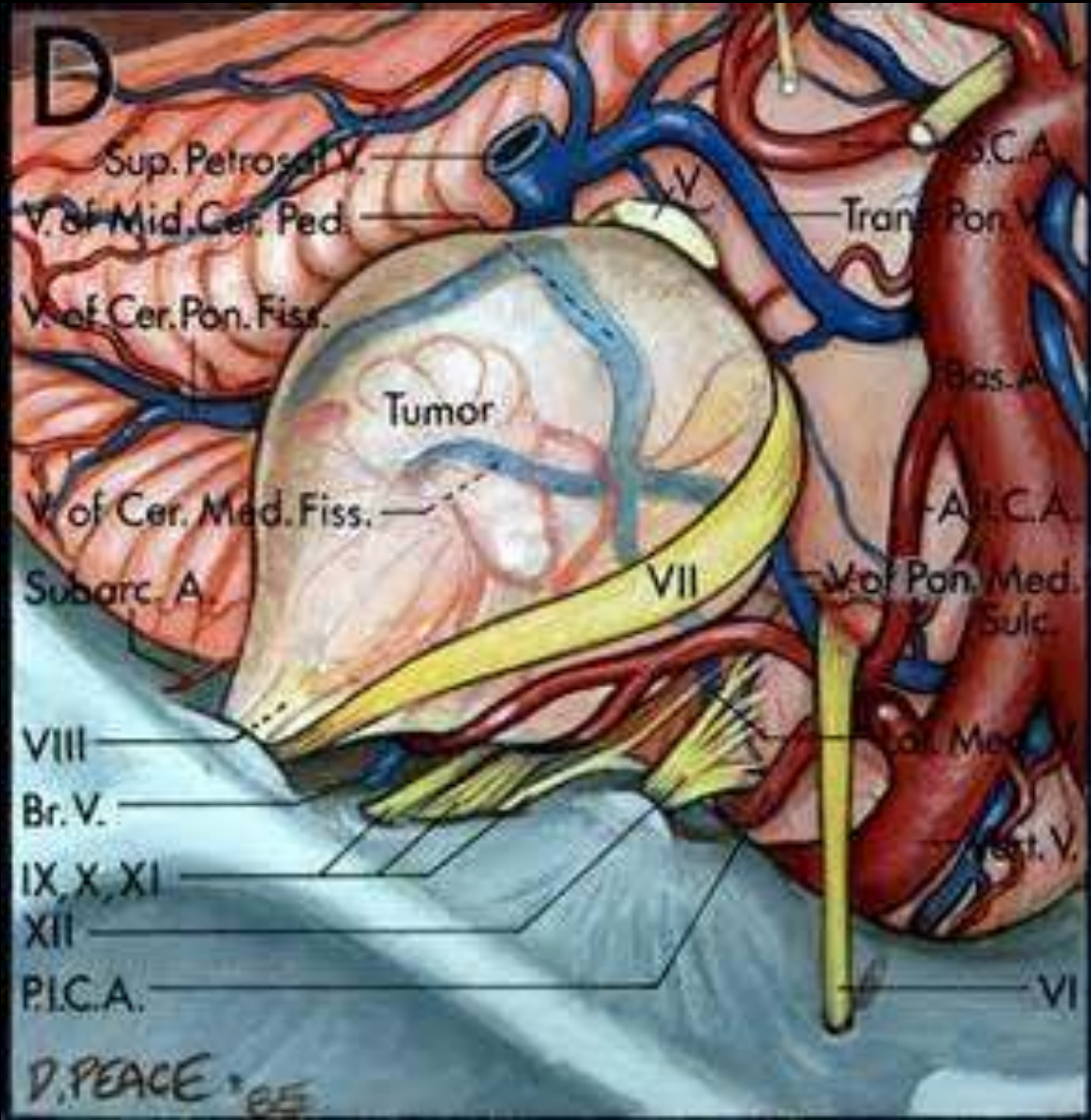
PROBLEM

Tumor location

Pressure on Facial
Nerve



**Injury of
facial nerve**



Growth patterns

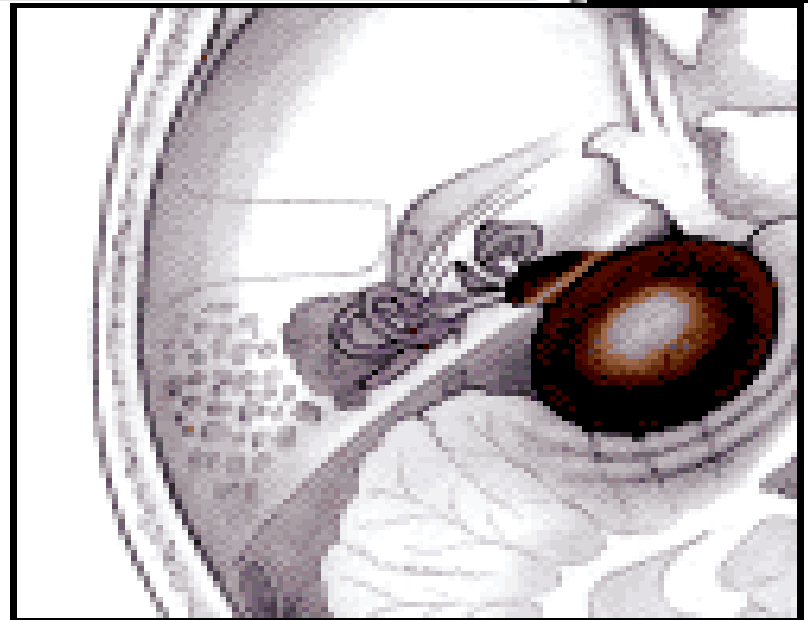
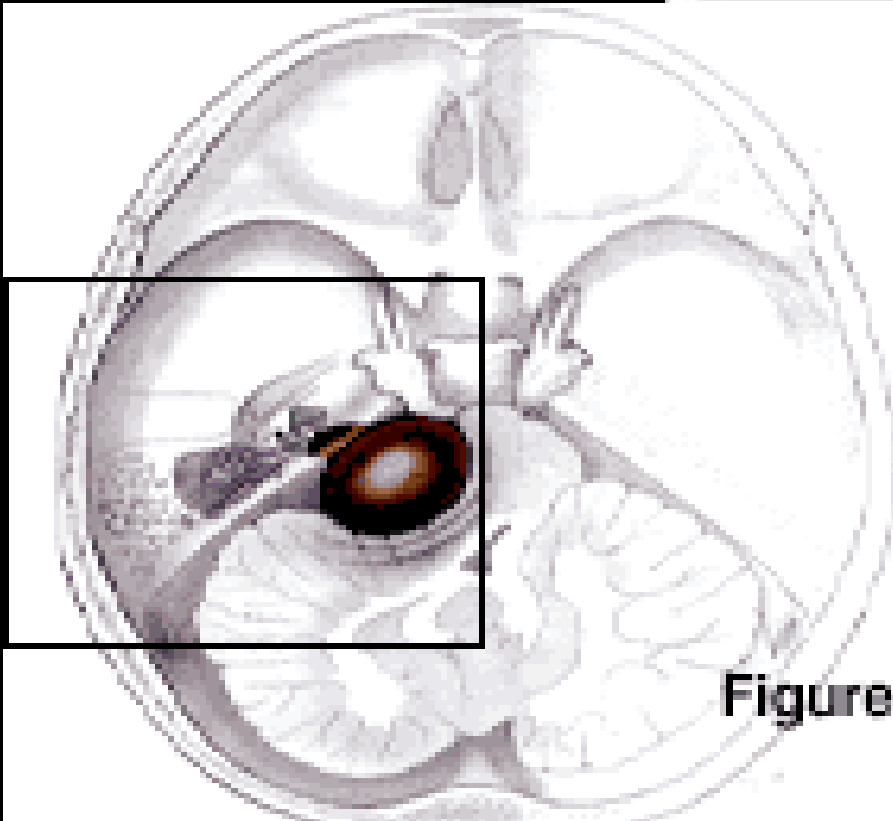
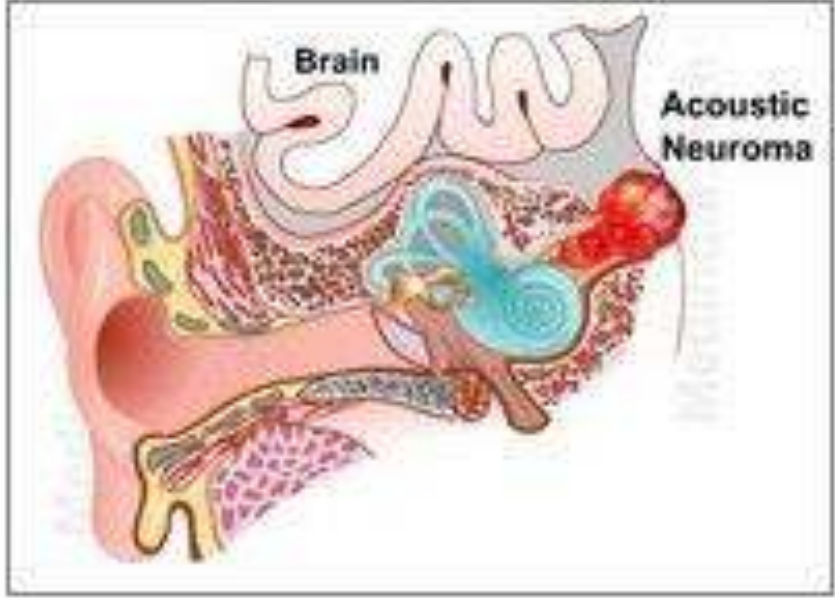


Figure 3. Brainstem compressive stage.

Paresis N. VII

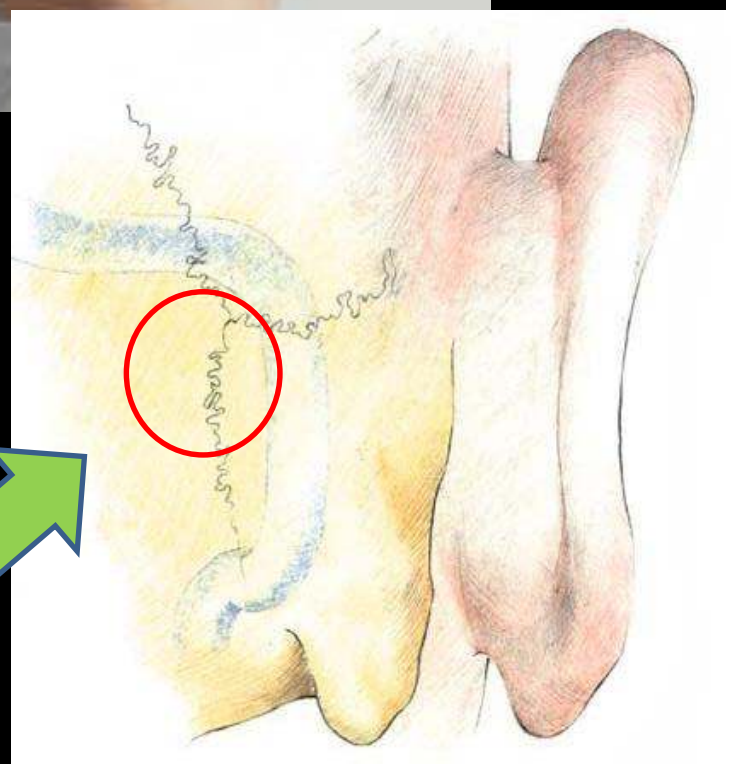
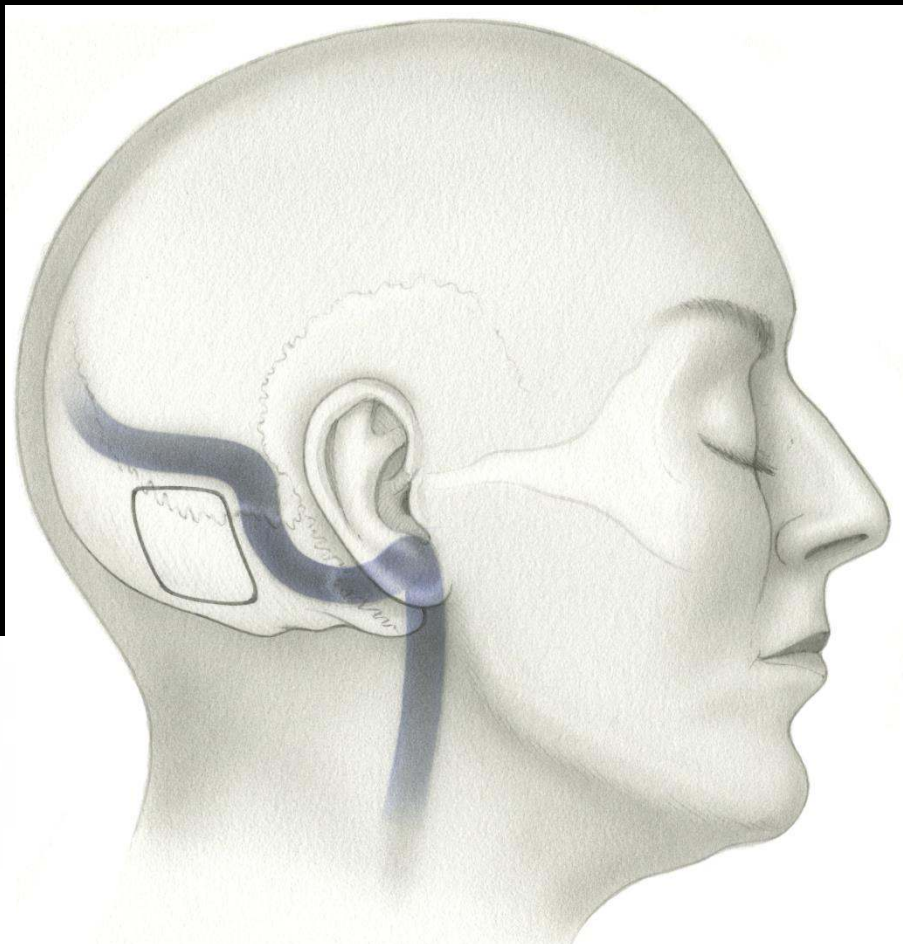


- ▣ **N.VII memiliki peran yang sangat kritis terhadap fungsi otot wajah**
- ▣ **cedera pada N.VII dapat sangat berat dan memberikan implikasi panjang secara fungsional, emosional, dan sosial sepanjang hidup pasien.**

Martin HC, Sethi J, Lang D, Neil-Dwyer G, Lutman ME, Yardley L: Patient-assessed outcomes after excision of acoustic neuroma: postoperative symptoms and quality of life. J Neurosurg 94:211–216, 2001

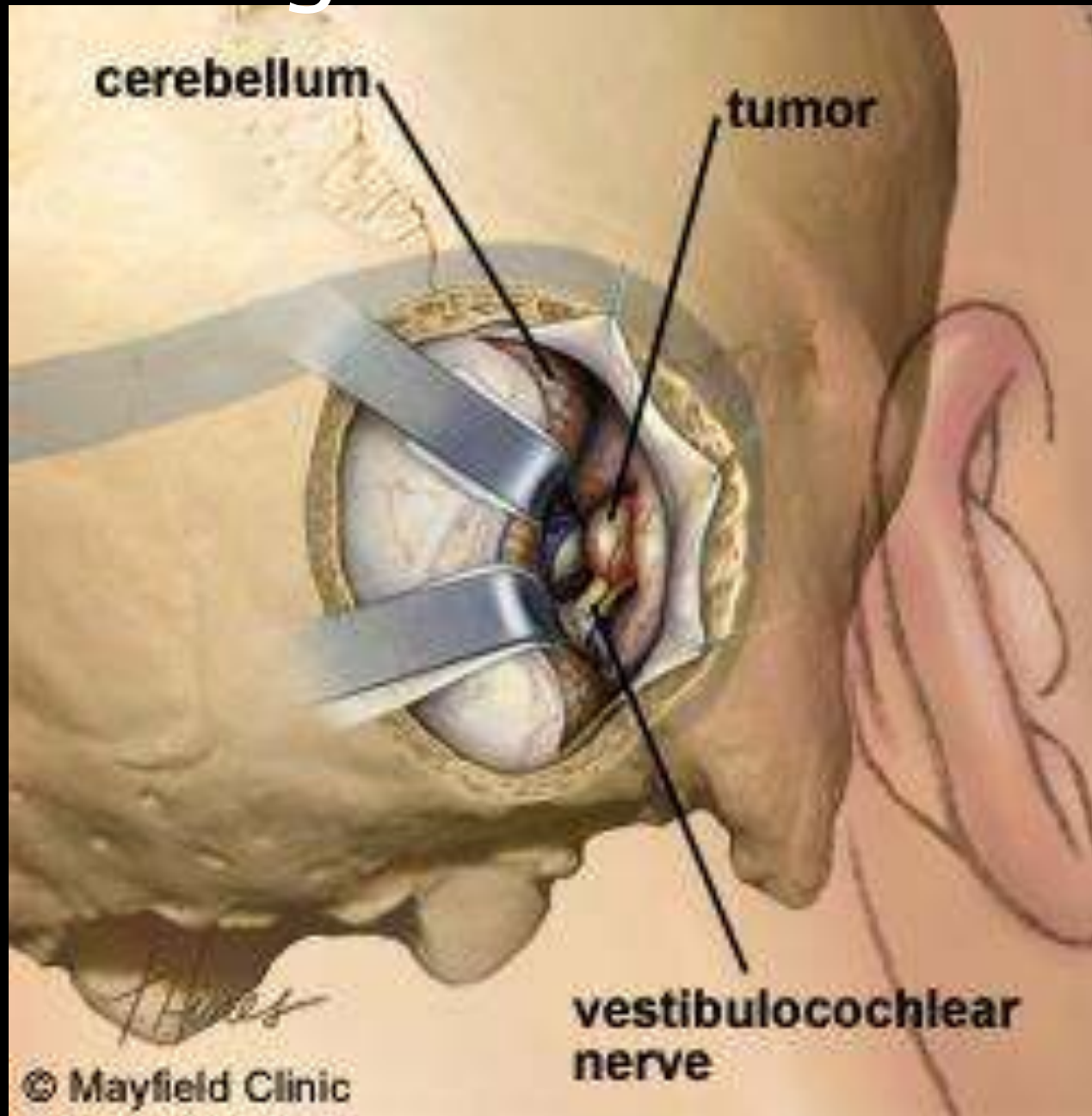
Vestibular Schwannoma

- ▣ Benign tumor → origin : vestibular nerve
- ▣ **CLINICAL TRIAD** :
 - Hearing loss (insidious & progressive)
 - Tinnitus
 - Dysequilibrium
- ▣ Tx option → depend on :
 - TUMOR SIZE
 - Growth
 - Hearing status

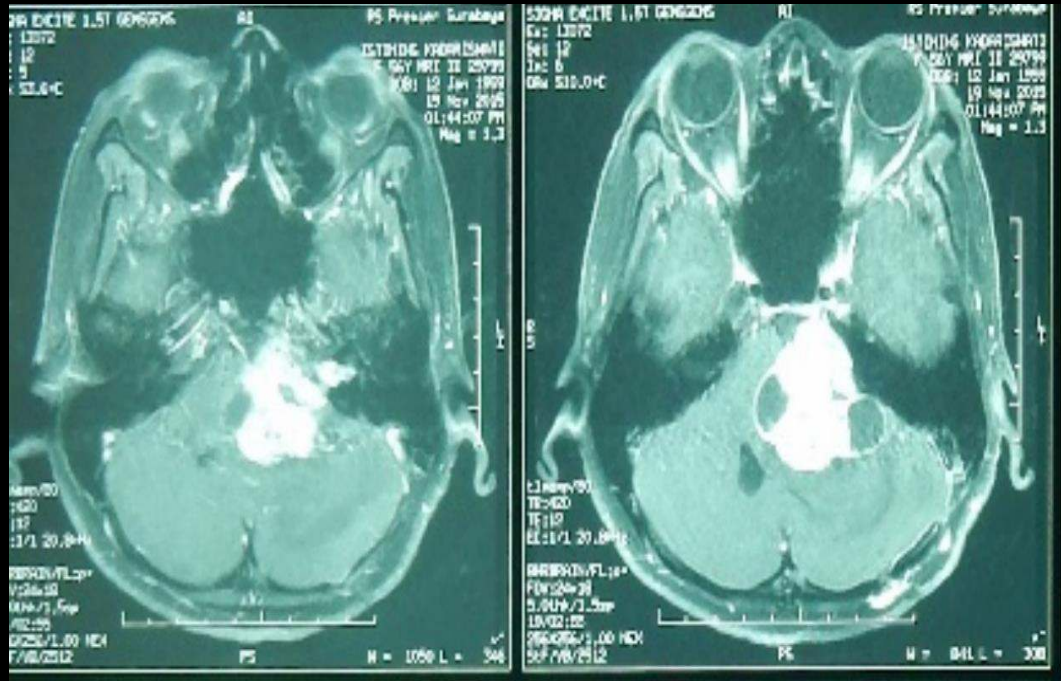


**Identification of
tranverse and sigmoid
sinus**

Retro-Sigmoid Craniotomy

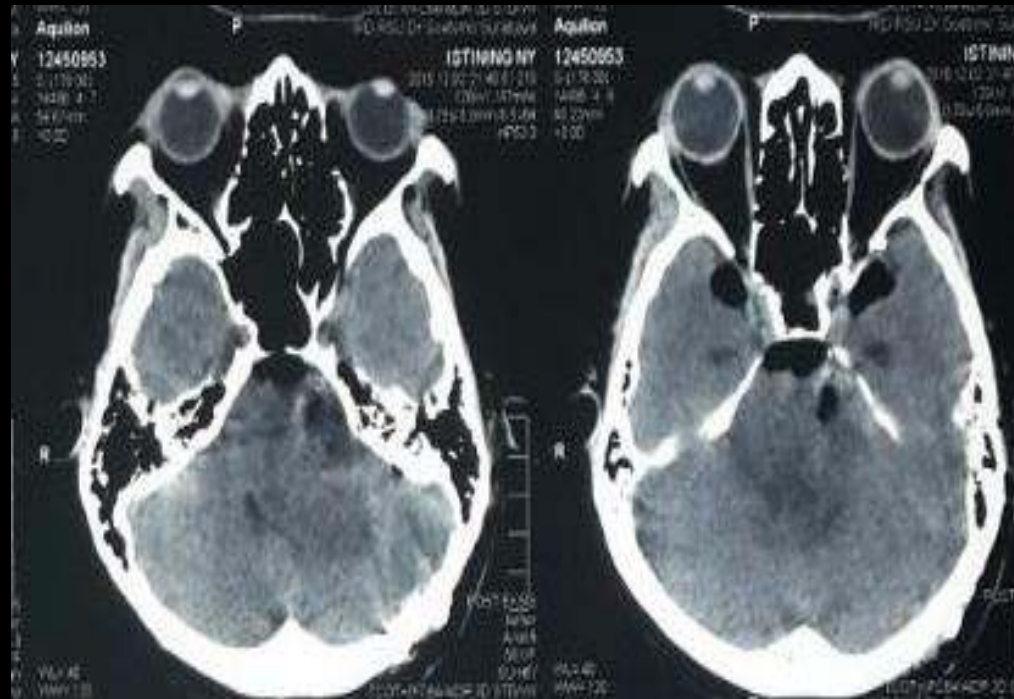


Cpa tumor (S) residif



Perempuan 56 th/ gangguan keseimbangan sejak 4 bulan, bicara pelo, telinga kiri berdenging, Riw Operasi tumor Juni 2015/ GCS 15 parese N VII (S), cerebellar sign (+)

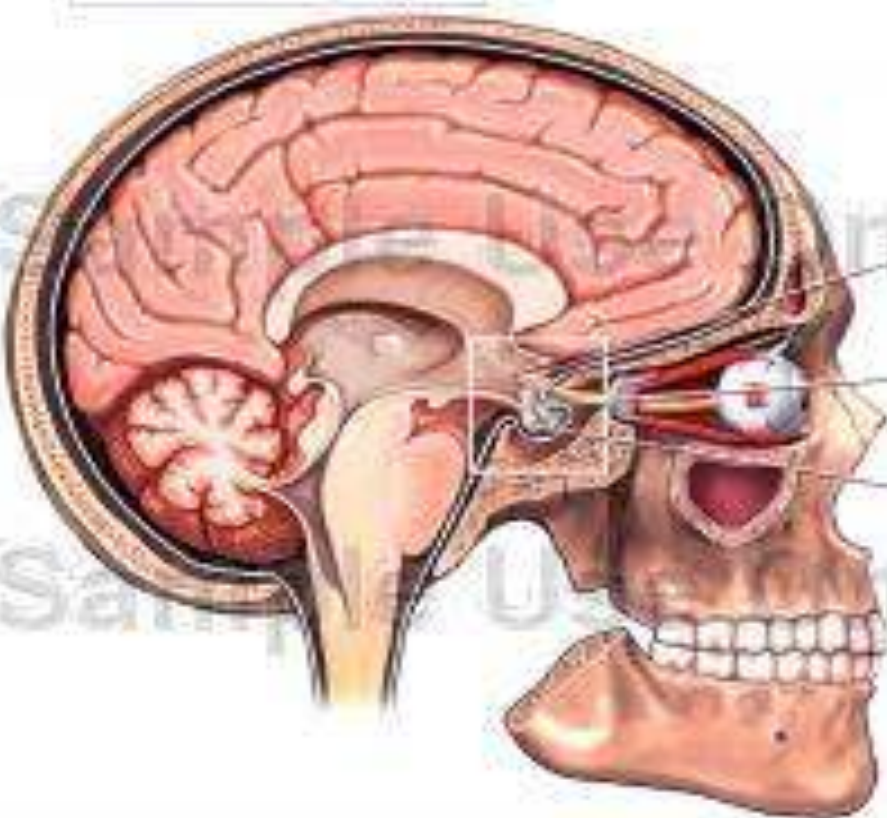
Post Op



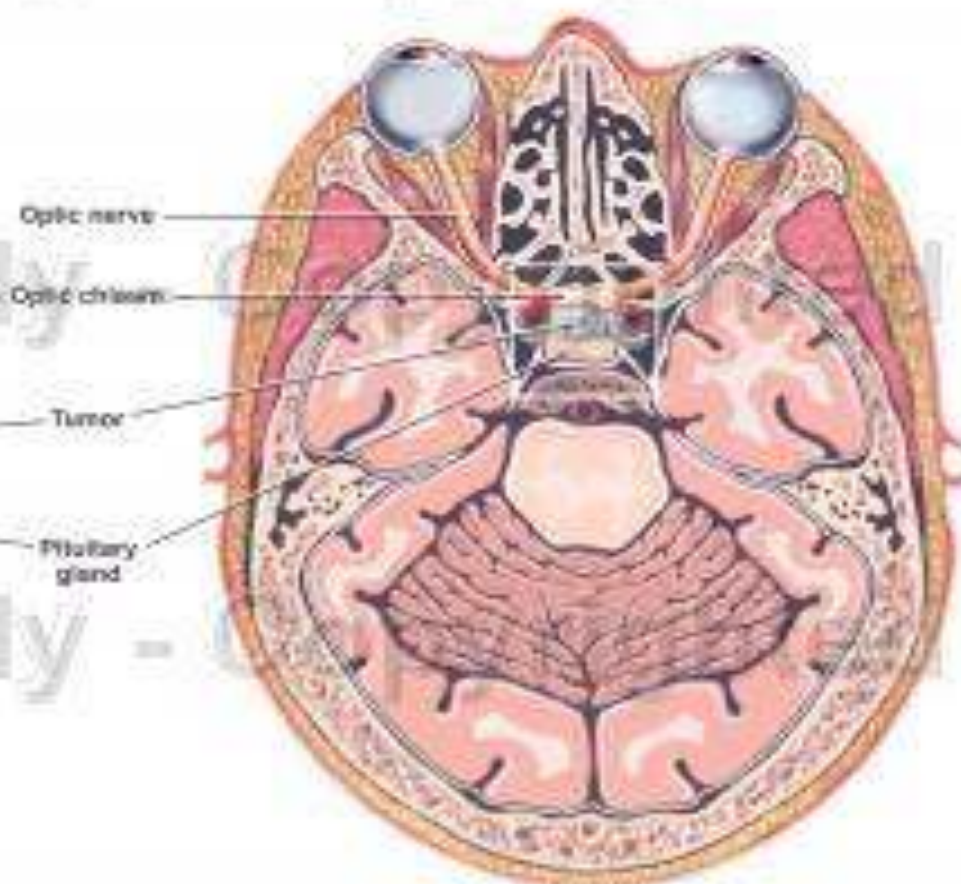
Pituitary Adenoma

Anatomy of the Brain Before Removal of Pituitary Tumor

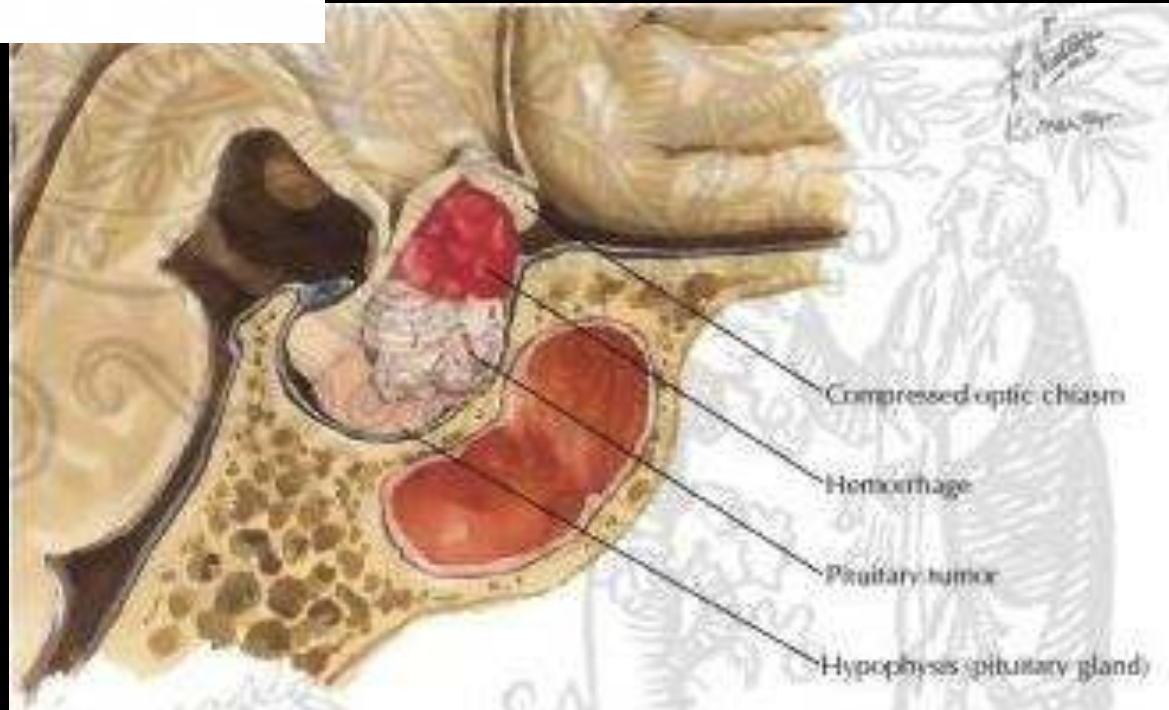
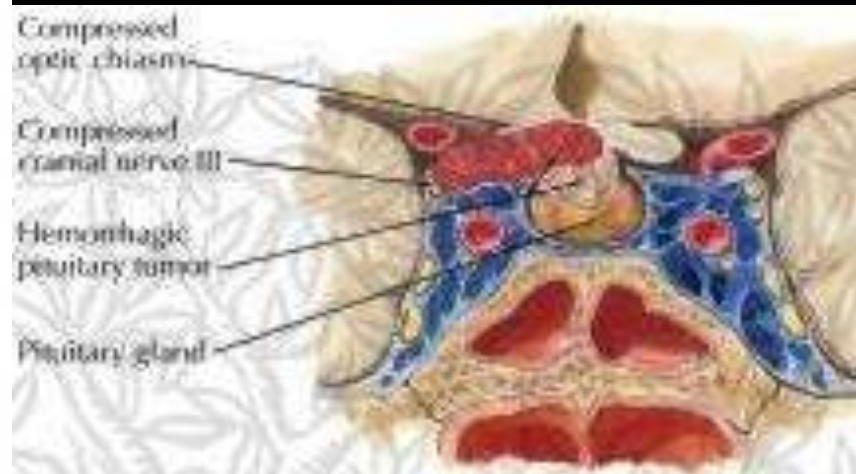
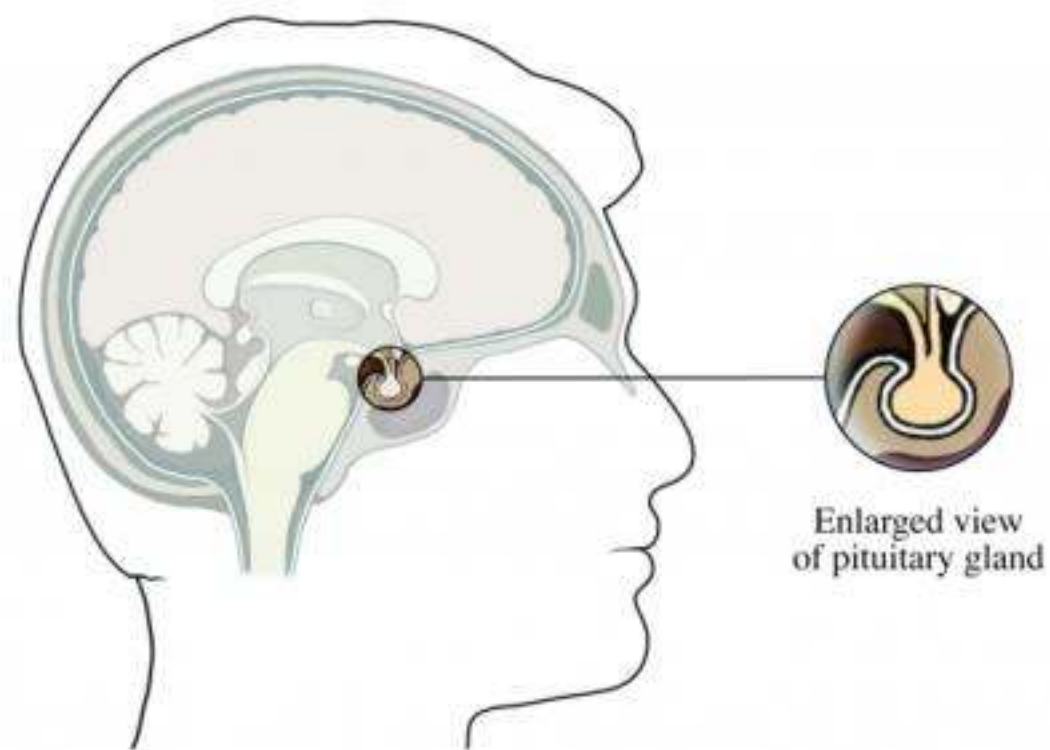
Large pituitary tumor compresses the optic chiasm, as well as invading the sphenoid sinus and cavernous sinuses bilaterally.



Lateral (cut-away) view



Superior (cross-sectional) view



Epidemiology

- ▣ Etiology is unknown
- ▣ Not associated with environmental factors
- ▣ 10-15% of all primary brain tumors
- ▣ 20-25% of pituitary glands at autopsy found to have adenomas
- ▣ **70% of adenomas are endocrinologically secreting**

Natural History

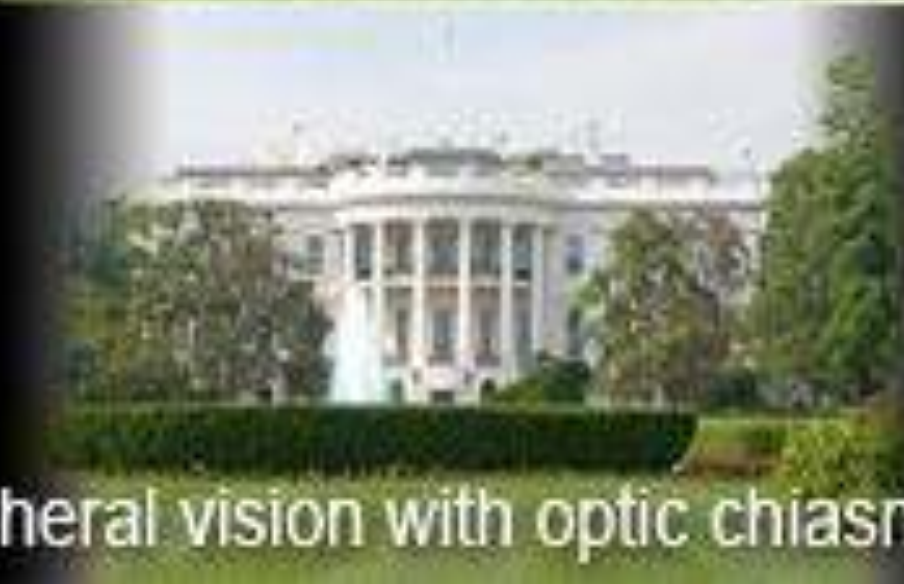
- ▣ Pituitary adenomas have long natural history
- ▣ Vary in size and direction of spread
- ▣ Microadenomas < 10 mm – may cause focal bulging
- ▣ Macroadenomas > 10 mm – cause problems due to mass effect

Clinical Presentation

- ▣ Most common are endocrine abnormalities →
 - hyper-/hyposecretion of ant. pituitary hormones
- ▣ Headache
- ▣ Vision changes
 - bitemporal hemianopsia and superior temporal defects



Penyempitan lapang pandang



TYPE

- ▣ **Endocrine-Active Pituitary Adenomas (Functioning Adenoma)**
- ▣ **Non-functioning Adenomas**

Endocrine-Active Pituitary Adenomas

- ▣ Prolactin → Amenorrhea, galactorrhea, impotence
- ▣ Growth hormone → Gigantism and acromegaly
- ▣ Corticotropin → Cushing's disease, Nelson's syndrome post adrenalectomy
- ▣ TSH → Hyperthyroidism

Non-functioning Adenomas

- ▣ 25-30 % of patients do not have classical hypersecretory syndromes
- ▣ **May grow to a large size before they are detected**
- ▣ Present due to mass effect :
 - **Visual deficits**
 - **Headache**
 - Hormone deficiency

Evaluation

- ▣ MRI
- ▣ **Visual field assessment**
- ▣ Endocrine evaluation
 - Tests of normal gonadal, thyroid, and adrenal function
 - Radioimmunoassays – for hormone levels

General Management

- ▣ Pituitary adenoma management is complex and is dictated by
 - size,
 - Symptoms,
 - character of tumor
- ▣ Treatment options require multiple modalities

General Management

- ▣ Multidisciplinary approach
- ▣ Goals:
 - Define tumor extent
 - Evaluate hormone activity
 - Remove tumor mass
 - Control hypersecretion
 - Correct endocrine deficiencies

Management of Pituitary Adenoma

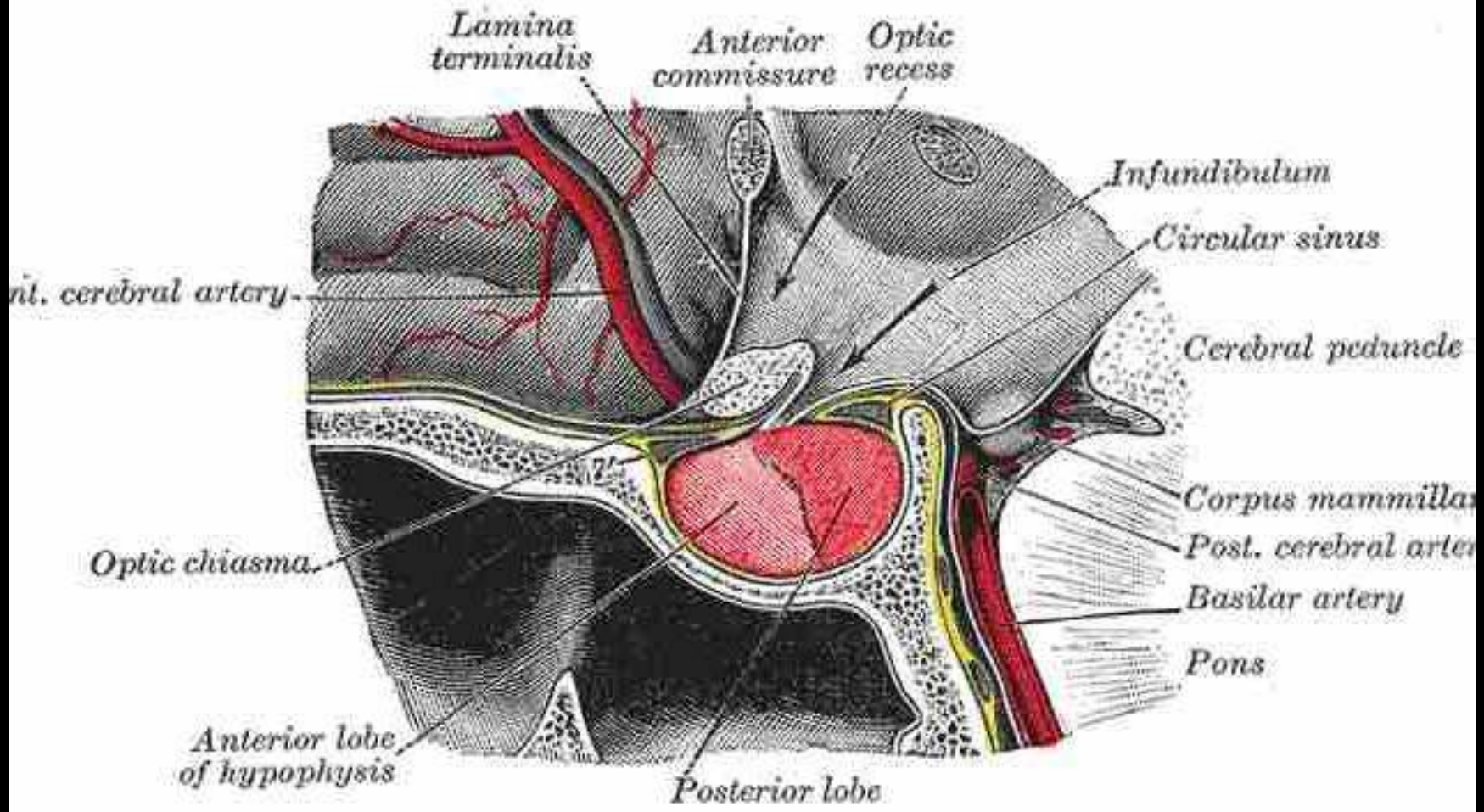
- ▣ Observation
- ▣ Pharmacotherapy
- ▣ Surgery
- ▣ Radiation Therapy (EBRT, SRS)

Indications for Surgery

- ▣ Surgery is the **first-line treatment** of symptomatic pituitary adenomas.
- ▣ Useful when medical or radiotherapy fails
- ▣ Surgery provides prompt relief from excess hormone secretion and mass effect.
- ▣ Indicated in **pituitary apoplexy** with compressive symptoms

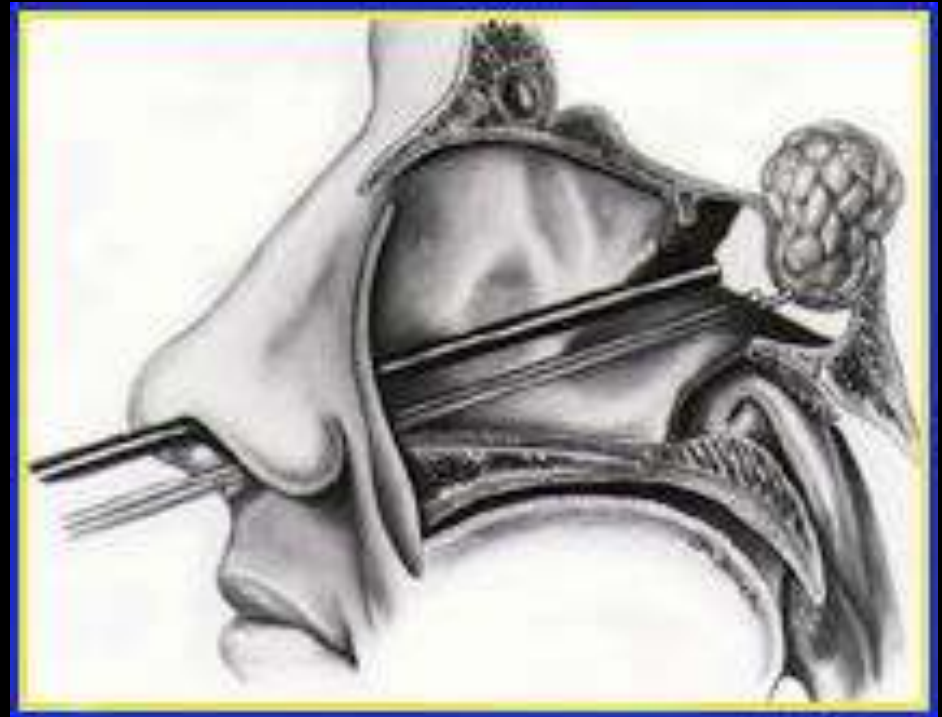
Pituitary apoplexy

- ▣ Sudden expansion of Pituitary adenoma (HEMORRHAGE or NECROSIS)
- ▣ Paroxysmal HEADACHE with endocrine deficiency
- ▣ Visual disturbance (usually ophthalmoplegia or visual loss)
- ▣ Increased ICP, Hypothalamic involvement, hydrocephalus
- ▣ Tx :
 - Immediate tx of pituitary function & endocrine evaluation
 - Decompression (Transphenoidal) → within 7 days

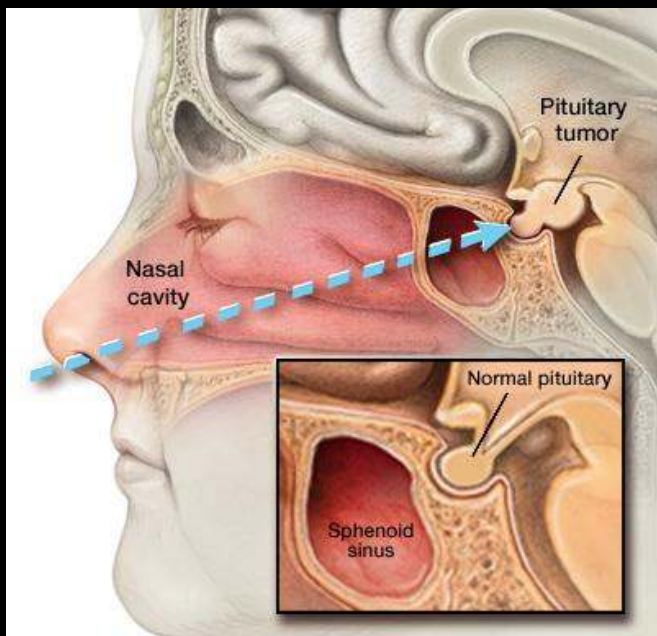


Pituitary Surgery

- ▣ Transsphenoidal approach: used for 95% of pituitary tumors
- ▣ Endonasal submucosal transseptal approach
- ▣ Septal Pushover/Direct Sphenoidotomy
- ▣ Endoscopic approach

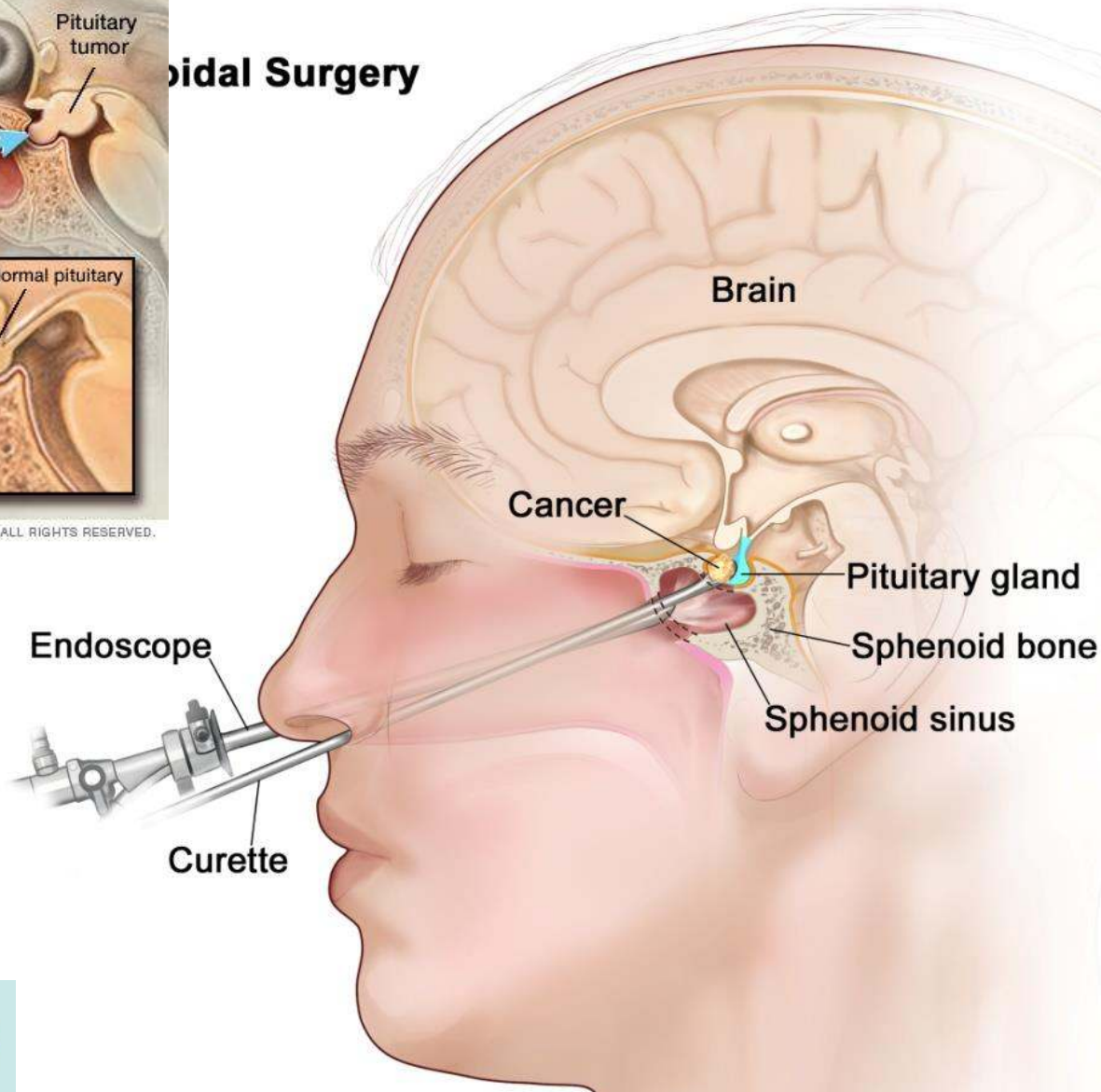


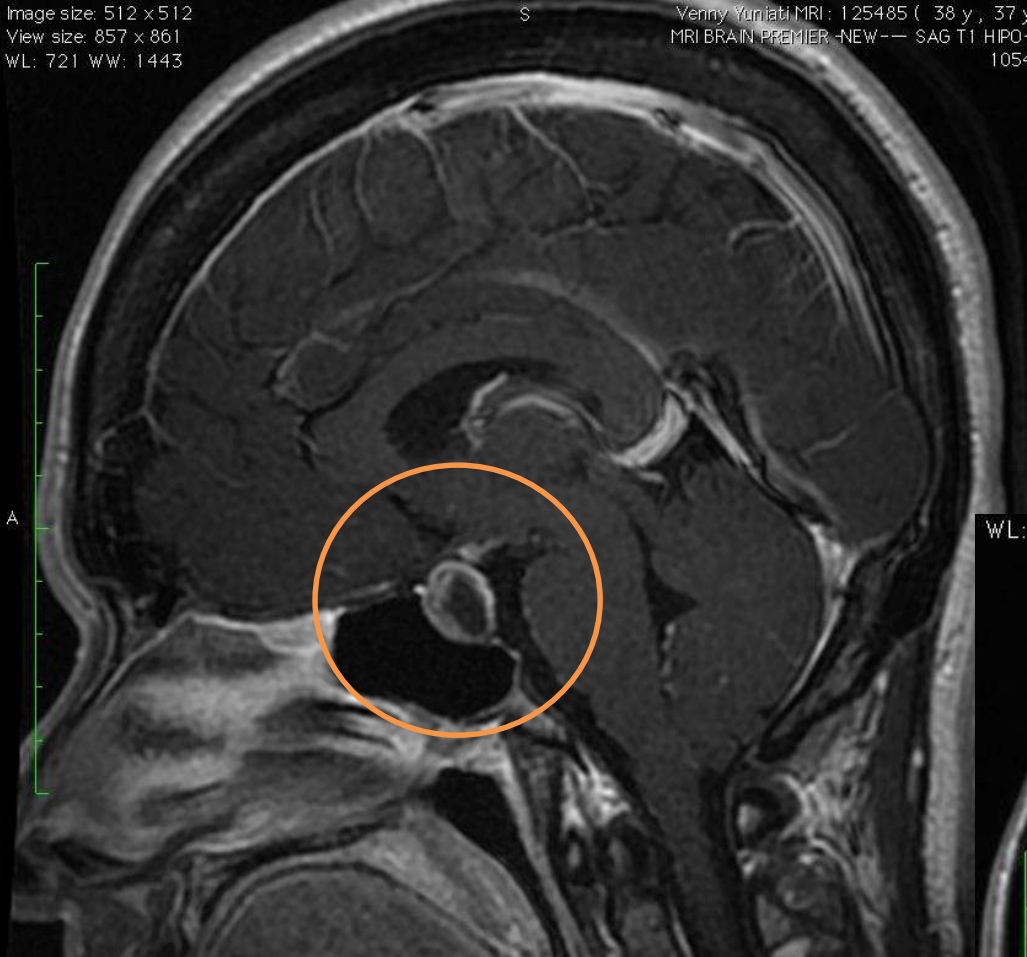
Endoskopi neurosurgery



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Sphenoidal Surgery



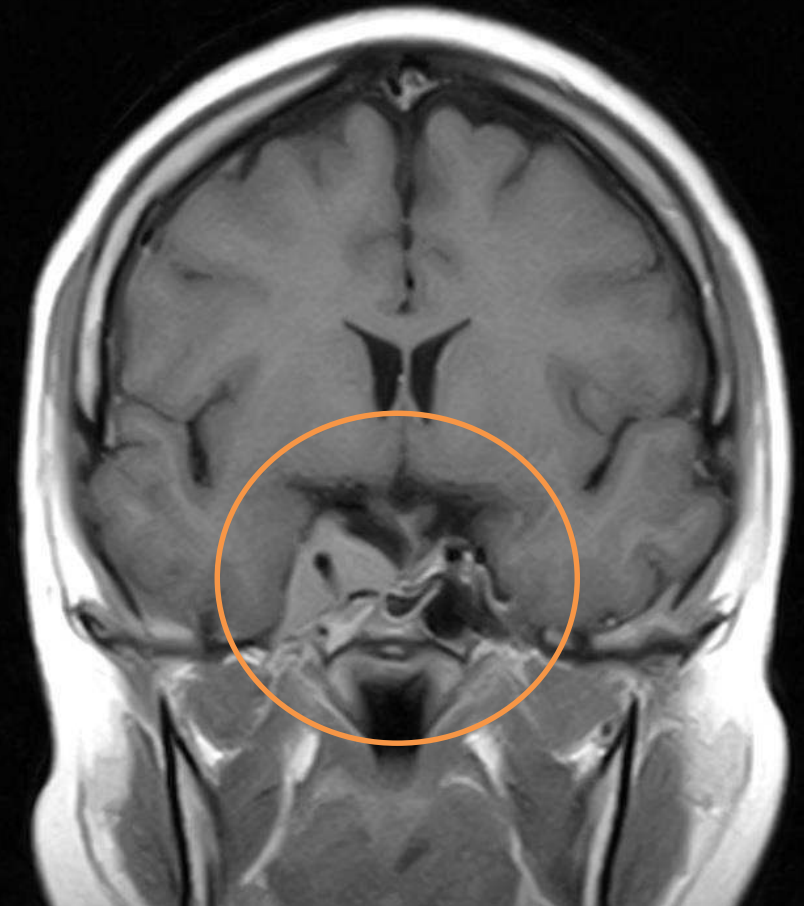
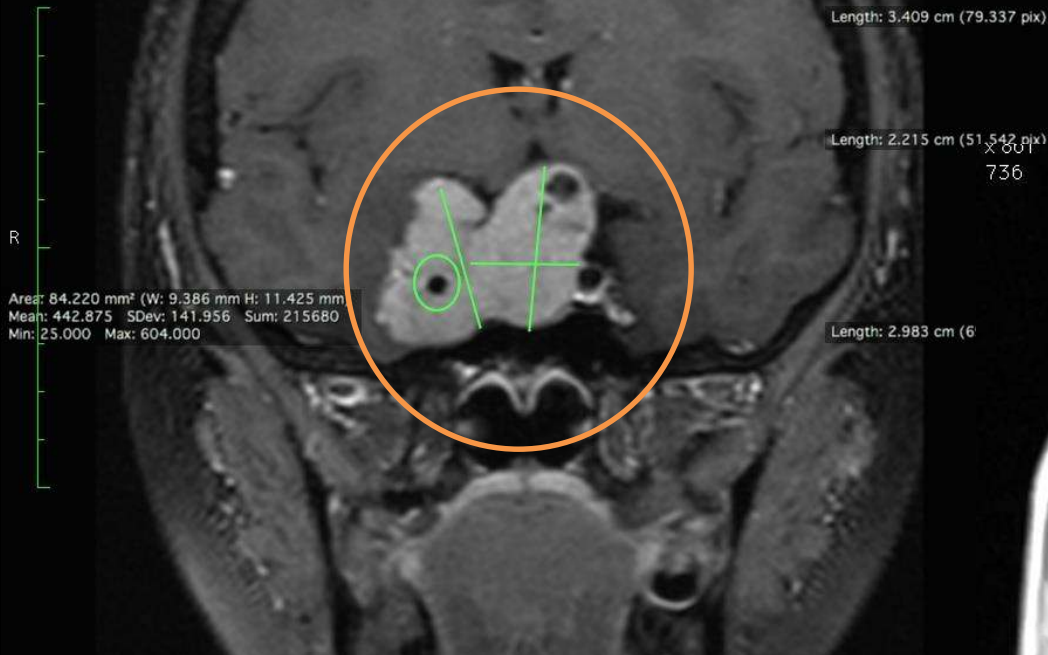


**Wanita, 34 tahun,
gangguan menstruasi,
tangan dan kaki makin membesar
Peningkatan Hormon
Pertumbuhan**

Image size: 512 x 512
View size: 903 x 861
WL: 478 WW: 976

SP

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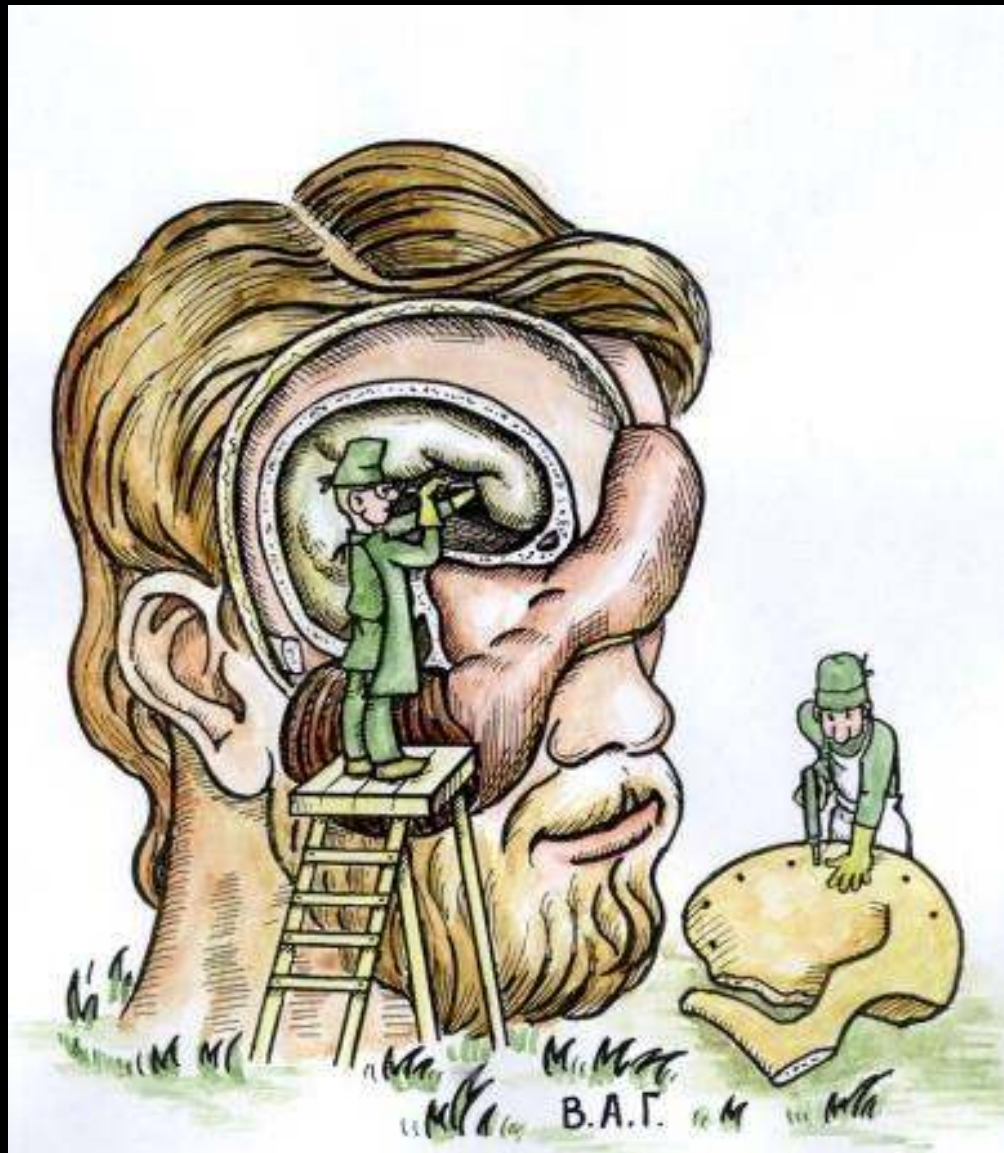


**Pria, 23 tahun,
Gangguan Penglihatan (lapang
pandang)
Gangguan ereksi
Peningkatan Hormon PROLAKTIN**

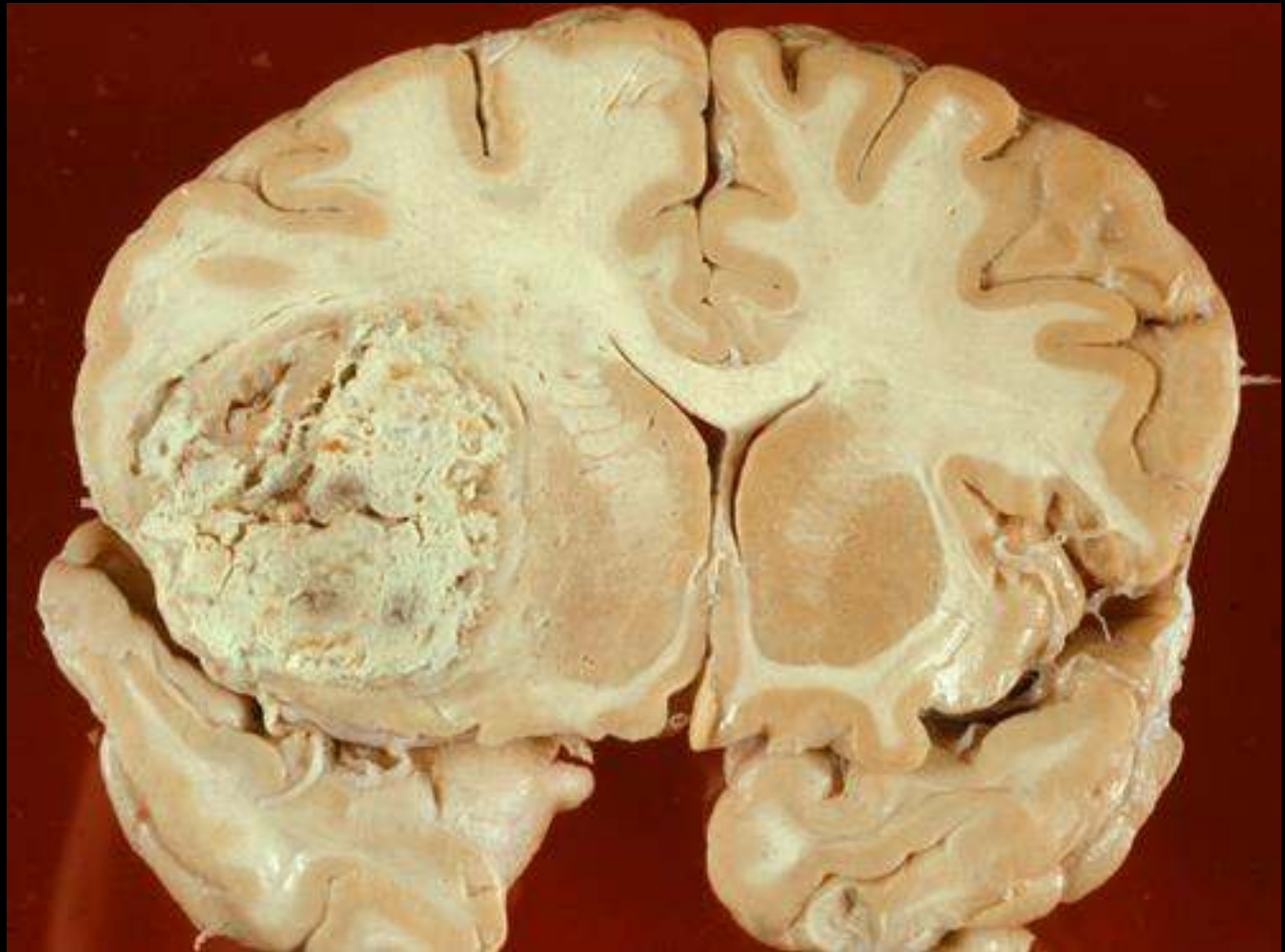
Radiation Therapy

- ▣ Reserved for patients with larger tumors and/or persistent hormonal hyperfunction despite surgical intervention
 - Conventional radiotherapy (EBRT)
 - Stereotactic Radio Surgery (SRS)
 - Gamma knife radiosurgery

Thank You



ASTROCYTOMA



WHO Classification

- ▣ Grade I (Pilocystic Astrocytoma)

- Most frequent brain tumors in children

- ▣ Grade II (low grade Astrocytoma)

- 25% of glioma & infiltrative

- ▣ Grade III (anaplastic Astrocytoma)

- Highly malignant, progress to GBM

- ▣ Grade IV (glioblastomamultiforme)

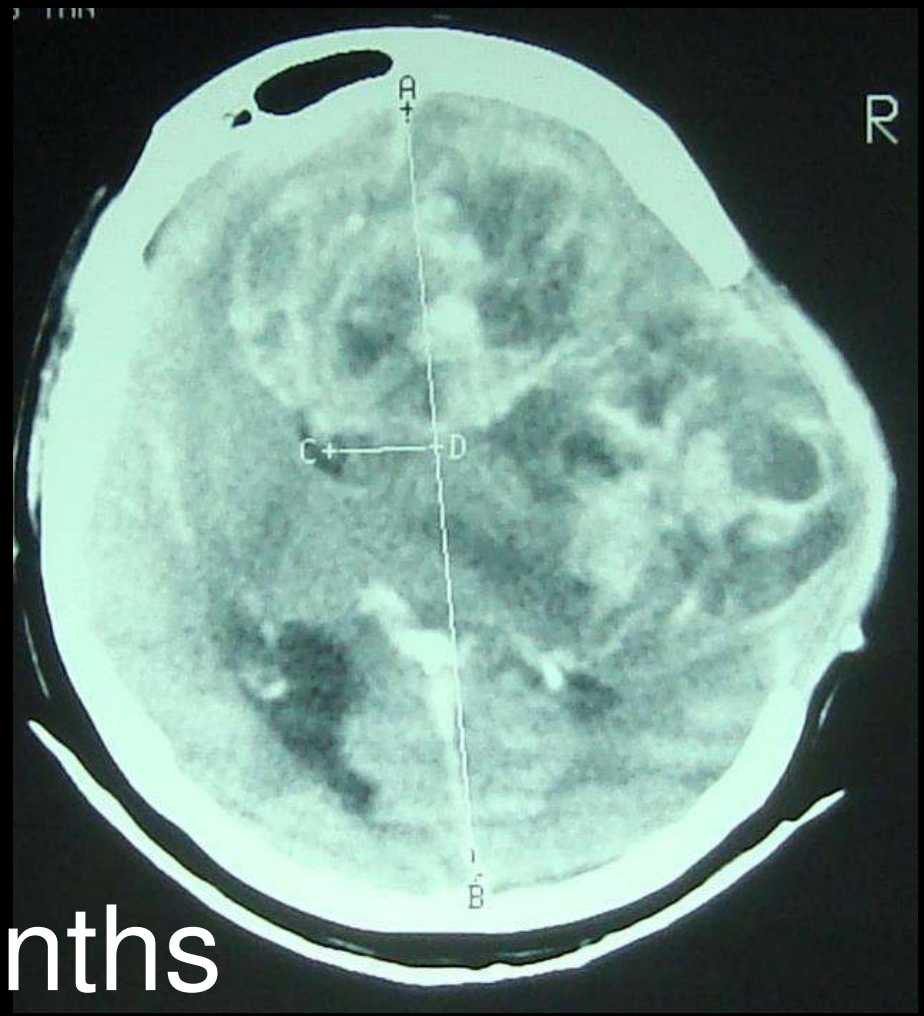
- Highly malignant, poor prognosis
- rapid spread to other region of the brain



Low grade



High grade



3 months

F/ 36 yo

Differentiated astrocytes or precursor cells

p53 mutation (>65%)
PDGF-A, *PDGFR-α*
overexpression (~60%)

Low grade astrocytoma

LOH 19q (~50%)
RB alteration (~25%)

Anaplastic astrocytoma

LOH 10q
PTEN mutation (5%)
DCC loss of expression (~50%)
PDGFR-α amplification (<10%)

Secondary glioblastoma

EGFR
amplification (~40%)
overexpression (~60%)

MDM2
amplification (<10%)
overexpression (~50%)

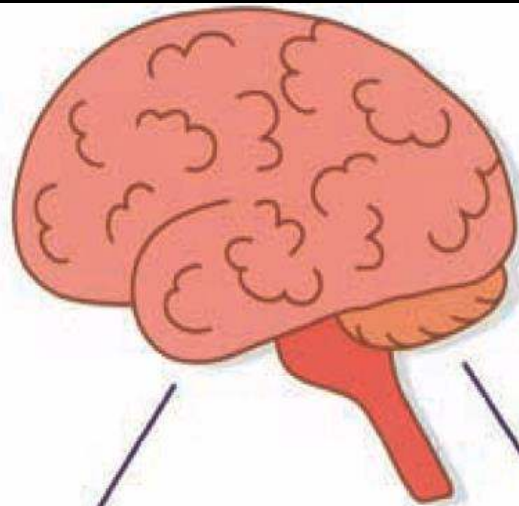
p16 deletion (30-40%)

LOH 10p and 10q
PTEN mutation (~30%)

RB alteration

**Primary glioblastoma
de novo**

Continuously mutating Brain Tumor



Clinical Features

- Low mitotic index
- Diffuse invasion
- High rate of transformation

**Low grade
Astrocytoma**

5-10 years

**Primary
Glioblastoma**

**Secondary
Glioblastoma**

Clinical Features

- Rapid proliferation
- Diffuse invasion
- Angiogenesis
- Cellular necrosis

Long term follow up

Treatment

● MULTIDISCIPLINARY

● OPTIONS :

● Surgery

● Radiation

● Chemotherapy

Major Prognostic Factors

- Age
- Karnofsky Performance Score
- Histological type
 - (oligodendroglial, mixed or astrocytic)
- Tumour grade
 - (WHO grade III or IV)

OTHER prognostic factors :

- **The extent of resection**
- The location and **volume of the tumour.**

Tx High grade glioma

- ▣ **Surgery is the initial recommended approach** in suspected high grade gliomas,
- ▣ **Goal of surgery**
 - debulking,
 - clinical improvement
 - pathologic diagnosis
 - **SET UP OTHER THERAPIES**

Resection vs Biopsy

- **Vuorinen et al (2003):**

“survival advantage for resection over biopsy”

- **Nomiya et al :**

retrospectively analyzed

the impact of the extent of resection exclusively
in 170 patients with Anaplastic Astrocytoma →
better survival.

Table 3: Evidence table for studies referred to in the systematic review for surgery

Authors ^{ref.} (Date)	Study Type	Tumour grade	Treatment allocation	Number of patients	Median Survival (Months)	P-value
Vuorinen et al. ²⁵ (2003)	RCT	III and IV	Resection ± RT ¹ Biopsy + RT	10 13	5.7 2.8	.0346
Lacroix et al ¹⁹ (2001)	Retrospective	IV	≥ 98% resection < 98% resection	197 219	13 8.8	0.0001
Shinoda et al ²⁶ (2001)	Retrospective	IV	GTR chemo ± RT < GTR chemo + RT	36 46	13 9.0	0.0002
Kreth et al. ⁵ (1999)	Retrospective	IV	Resection ± RT Biopsy + RT	126 99	8.8 7.8	0.09
Mohan et al. ³² (1998)	Retrospective	IV	Biopsy only Biopsy and radiation STR ² and radiation GTR ¹ and radiation	25 28 42 07	1.2 5.1 7.2 17.3	0.0001
Kowalczyk et al. ²⁰ (1997)	Retrospective	III and IV	GTR chemo ± RT STR chemo ± RT Biopsy chemo + RT	29 33 13	27.6 34 13	0.039
Kiwit et al ²⁷ (1996)	Please complete	IV	GTR Biopsy	40 40	10 6	<0.05
Slotman et al ²⁴ (1996)	Prospective	IV	>75% resection ± RT <75% resection + RT	20 8	10 7	<0.05
Quigly et al ¹⁸ (1995)	Retrospective	III and IV	GTR STR Biopsy	9 31 23	27 11 10	<0.05

Table 3 (contd.): Evidence table for studies referred to in the systematic review for surgery

Authors (Date)	Study Type	Tumour grade	Treatment allocation	Number of patients	Median Survival (Months)	P-value
Nitta et al. ²⁸ (1995)	Prospective	IV	GTR chemo ± RT STR chemo ± RT PR ⁺ chemo ± RT	14 27 27	53 14 14	<0.01
Kelly et al. ²¹ (1994)	Prospective	IV	SVR ⁵ ± RT SB ⁵ ± RT	40 88	6.3 ⁷ 3.6	0.008
Simpson et al. ³³ (1992)	Prospective	IV	Partial resection ± chemo ± RT Biopsy + chemo ± RT Total resection ± chemo ± RT	413 107 125	10.4 6.6 11.3	<0.001 ⁸ <0.0001 ⁹
Laws et al. ³⁷ (2003)	Prospective	IV III	Biopsy Craniotomy Biopsy Craniotomy	84 329 41 106	4.9 10.57 12.16 20.3	<0.0001 <0.0001
Stummer et al. ³⁵ (2006)	RCT	High grade gliomas	Surgery with 5-aminolevulinic acid Surgery with white light	139 131	5.1 ¹⁰ 3.6	<0.0003
Stummer et al. ³⁴ (2008)	RCT	IV	Partial resection Complete resection	121 122	11.8 16.9	<0.0001

Surgery

- ▣ Surgical resection as the main treatment for
 - alleviation of symptoms and
 - prolongation of life
- ▣ **Most studies report a survival advantage for resection over biopsy**
 - Alberta Cancer Board, Clinical Practical Guideline, Management of Glioblastoma Multiforme, 2009
- ▣ **Recommendation:**
gross total resection, **when safe**,
in high grade gliomas (WHO grade III and IV).

Adjuvant Chemo- Radiation Tx

“the standard of care following surgery”

1. surgery
2. followed by adjuvant combined Temozolomide-based chemotherapy and radiation →
“Concomitant Treatment”
3. followed by adjuvant Temozolomide for a total of six to 12 cycles.

Radiation Tx

- ▣ External beam radiation therapy will be given in standard fractionation to a maximum total dose of 59.4 - 60 Gy using 3D conformal planning techniques.
 - **TUMOR PLUS 1 – 2 CM FIELD**
- ▣ The volume treated should be partial brain irradiation and not whole brain irradiation
- ▣ No evidence to recommend a total dose >60 Gy in standard fractionation

Elderly Px

- ▣ Elderly patients (≥ 60 years old) with a poor performance status (KPS ≤ 70), \rightarrow adjuvant radiation tx alone, with or without tissue diagnosis
- ▣ Radiation Tx \rightarrow abbreviated to 40 Gy in 15 fractions

cases

1. Male, 56 y.o, concomitant dose 160 mg for 45 days, adjuvant dose 380 mg (6 cycles).
2. Male, 35 y.o, concomitant dose 160 mg for 45 days, adjuvant dose 380 mg (6 cycles).
3. Male, 35 y.o, concomitant dose 140 mg for 45 days, adjuvant dose 320 mg (6 cycles).
4. **Female**, 48 y.o, concomitant dose 120 mg for 45 days, adjuvant dose 280 mg (6 cycles).
5. Male, 55 y.o, concomitant dose 160 mg for 45 days, adjuvant dose 380 mg (6 cycles).
6. Male, 28 y.o, concomitant dose 140 mg for 45 days, adjuvant dose 360 mg (5th cycle).
7. Male, 45 y.o, concomitant dose 160 mg for 45 days, adjuvant dose 360 mg (4rd cycles).
8. Male, 20 y.o, concomitant dose 160 mg for 45 days, adjuvant dose 400 mg (3rd cycle).
9. Male, 10 y.o, adjuvant dose 220 mg (6 cycles).
10. Male, 54 y.o, adjuvant dose 280 mg (3rd cycle).
11. **Female**, 18 y.o, adjuvant dose 340 mg (1st cycle).

Result

- ▣ No major side effect that need medical tx
- ▣ Outcome : → follow up

Outcome

- ▣ Need long term follow up
- ▣ Future : Indonesia Brain Tumor Registry

THANK YOU