



EPILEPSY

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Objectives

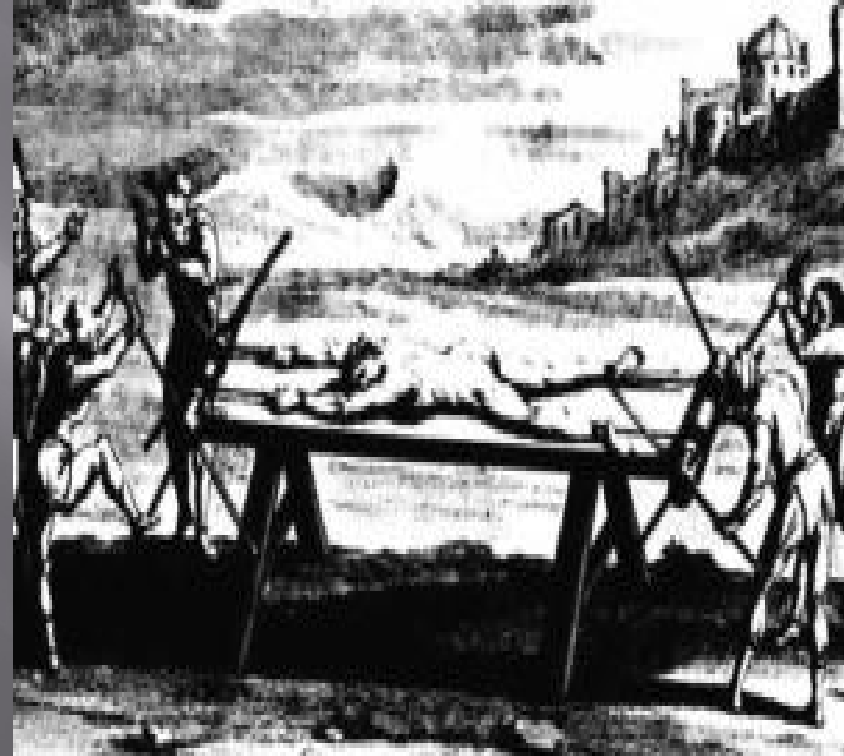
- By the end of this lecture you should be able to know:
 - Definition of epileptic seizure, provoked seizure and epilepsy.
 - Status epilepticus.
 - Frequent causes of seizure and risk factors.
 - Triggers of seizures in epileptic patient.
 - Epilepsy classification and seizure semiology.
 - DDX of SZ
 - Seizure vs syncope
 - Approach to seizure disorder (Hx, Ex, inx)
 - Medical and surgical management of epilepsy.
 - How to select antiepileptic medications.
 - When to stop antiepileptic medications.

History

- ▣ One of the earliest descriptions of a secondarily generalized tonic-clonic seizure was recorded over 3000 years ago in Mesopotamia.
- ▣ The seizure was attributed to the god of the moon.
- ▣ Epileptic seizures were described in ancient cultures, including those of China, Egypt, and India.

History

- ▣ Hippocrates wrote the first book about epilepsy almost 2500 years ago.
- ▣ He rejected ideas regarding the divine etiology of epilepsy and concluded that the cause was excessive phlegm that caused abnormal brain consistency.





What is epilepsy?

- ▣ **Epileptic seizure:** transient occurrence of signs and symptoms of sudden changes in neurological function due to abnormal excessive ,synchronous discharge of cortical neurons ..
- ▣ **Provoked seizures:** is occur in the setting of acute medical and neurological illnesses in people with no prior history of seizures
- ▣ **Epilepsy:** recurrent (two or more) unprovoked seizures.
- ▣ Seizure is a symptom of epilepsy.

Status Epilepticus

- ▣ **Status epilepticus (SE):** defined as recurrent convulsions that last for more than 20 minutes and are interrupted by only brief periods of partial relief.
- ▣ **(SE):** is a serious, potentially life-threatening.
- ▣ Any type of seizure can lead to SE, the most serious form of status epilepticus is the generalized tonic-clonic type.



Epidemiology

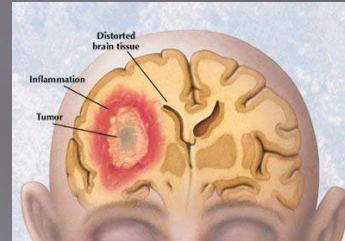


Epidemiology and course

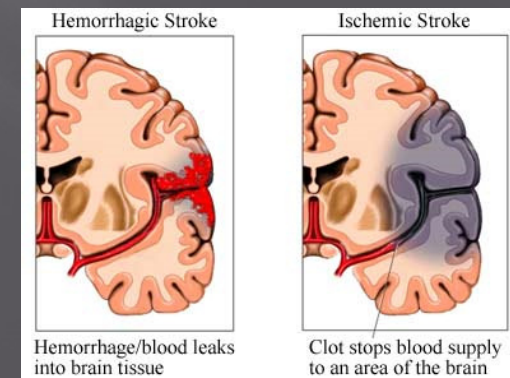
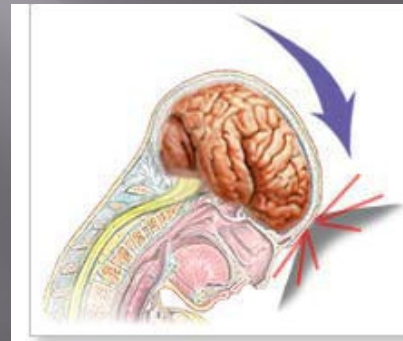
- **5%** of the population suffer a single sz at some time
- **0.5-1%** of the population have recurrent sz = EPILEPSY
- **70%** = well controlled with drugs (prolonged remissions)
- **30%** epilepsy at least resistant to drug treatments = INTRACTABL EPILEPSY.

Risk Factors for Epilepsy

- ▣ Febrile convulsion
- ▣ Perinatal insult
- ▣ CNS infection
- ▣ CNS mass lesion
- ▣ Family history of epilepsy
- ▣ Head injury
- ▣ Abnormal gestation or delivery
- ▣ Developmental delay
- ▣ Stroke (ischemic or hemorrhagic)

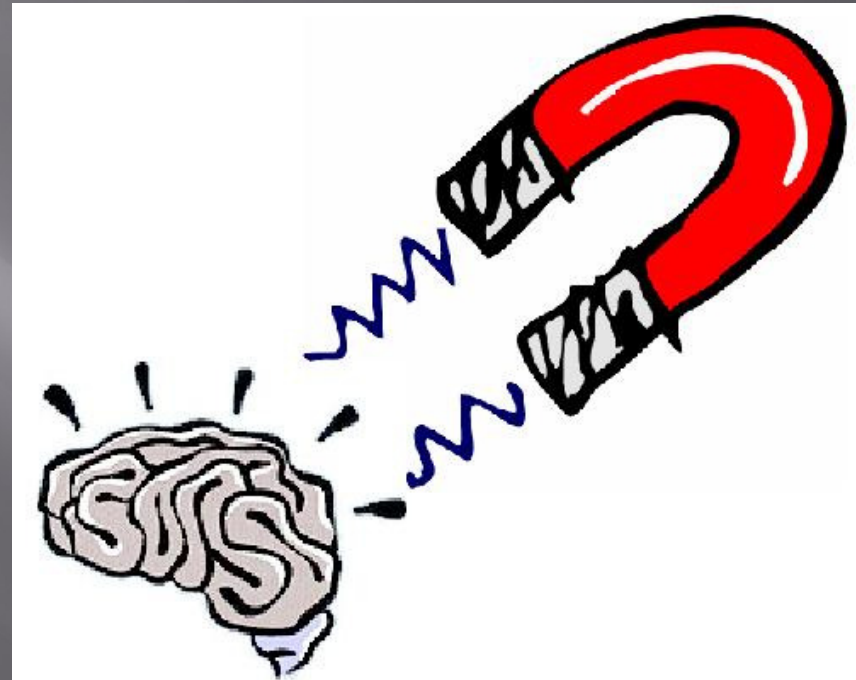


65%
unknown
cause



Triggers for seizure:

- ▣ Poor compliance
- ▣ Sleep deprivation
- ▣ Stress
- ▣ Alcohol
- ▣ Infection
- ▣ Menstrual cycle



DANGER



Status Epilepticus

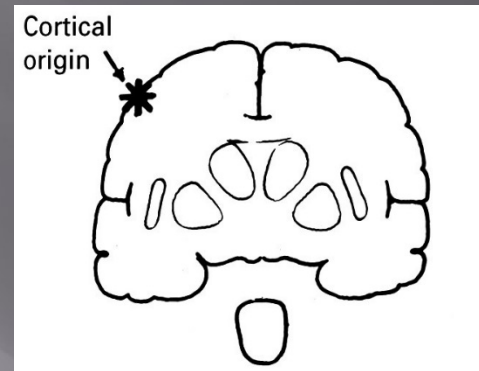
**SUDDEN
DEATH**



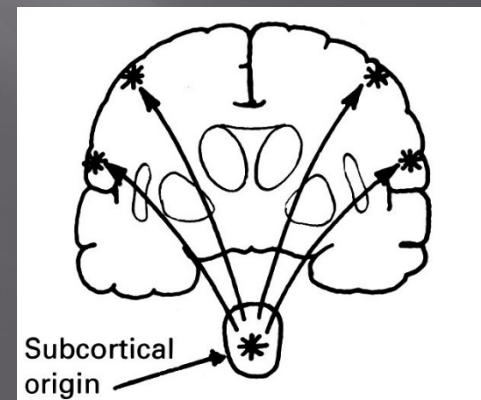
Seizure Classification

Epilepsy - Classification

- ▣ **Focal seizures** – *account for 80% of adult epilepsies*
 - Simple partial seizures
 - Complex partial seizures
 - Partial seizures secondarily generalised



- ▣ **Generalised seizures**
- ▣ **Unclassified seizures**



NEW ILAE Classification of seizures

Generalized seizures

Tonic-clonic (in any combination)

Absence

Typical

Atypical

Absence with special features

Myoclonic absence

Eyelid myoclonia

Myoclonic

Myoclonic atonic

Myoclonic tonic

Clonic

Tonic

Atonic

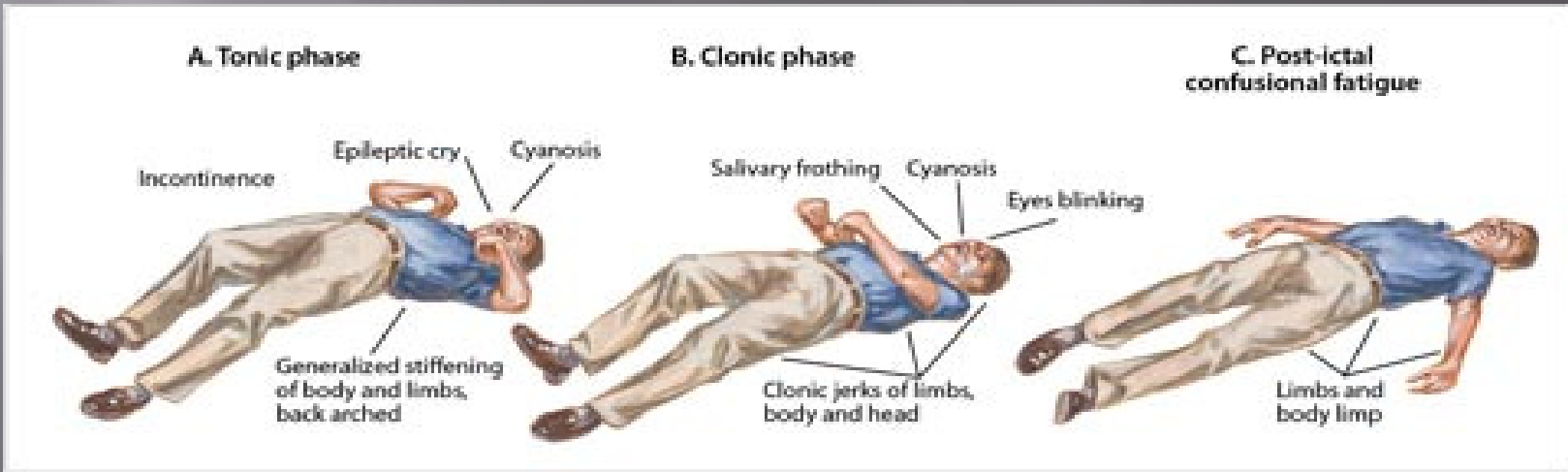
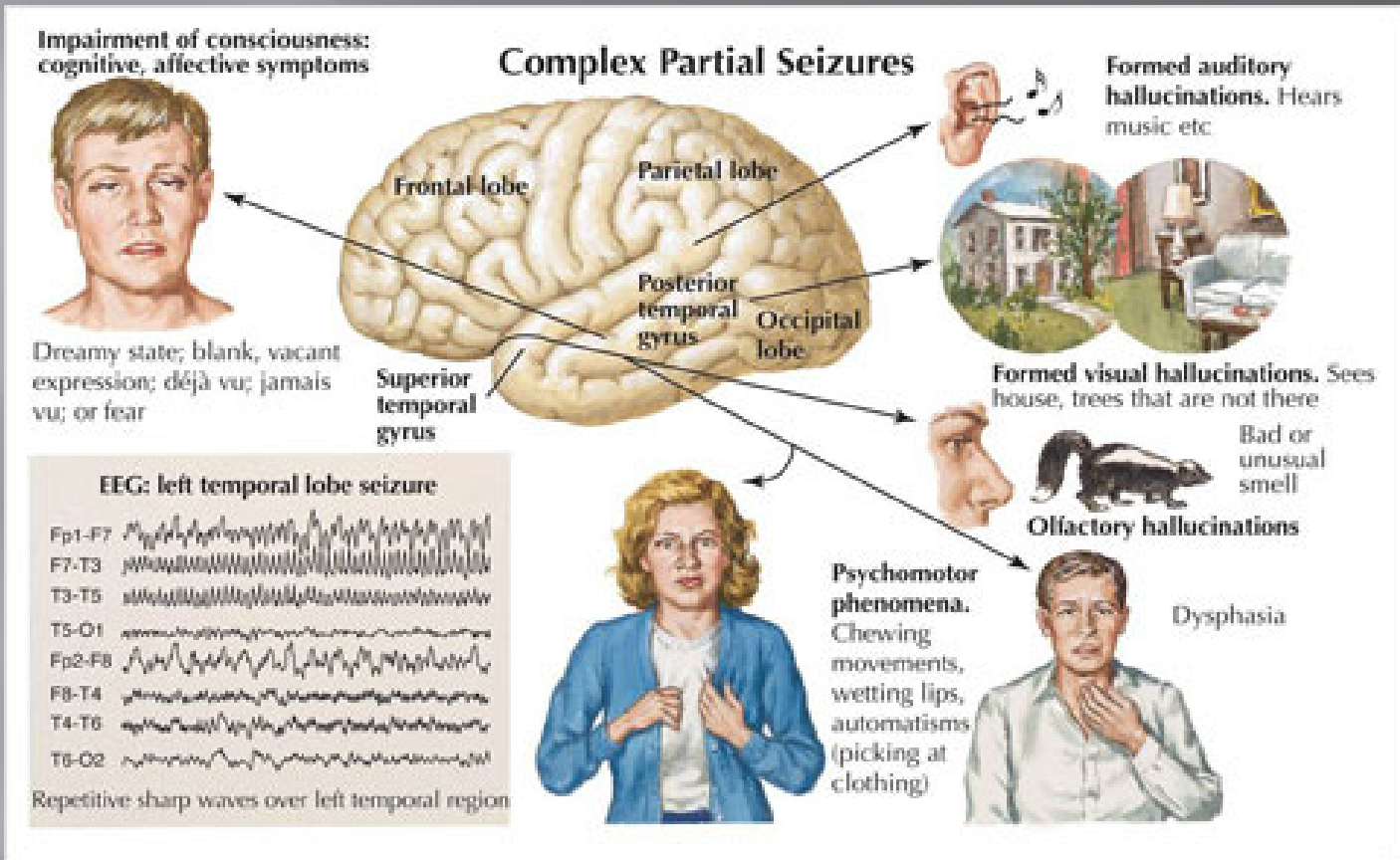
Focal seizures

Unknown

Epileptic spasms

Seizure Semiology





VEDIO













Summary of typical signs on video EEG

Typical EEG sign	Localizes to
Oral automatisms	Temporal lobe
Hypermotor automatisms	Frontal lobe
Manual picking automatisms	Temporal lobe
Visual hallucinations	Occipital lobe
Auditory hallucinations	Temporal neocortex (Heschl's Gyrus)
Olfactory hallucinations	Mesial temporal lobe
Nystagmus, eye blinking, eye pulling sensation	Occipital lobe
Ictal amaurosis	Occipital lobe
Tonic arm elevation	Supplementary motor area
Epigastric aura	Temporal lobe
Throat tightening sensation	Insula
Ictal pain	Parietal lobe
Somatosensory sensations	Postcentral gyrus or supplementary motor area
Clonic activity	Precentral gyrus
De-ja vu or jamais vu aura	Mesial temporal lobe
Fear	Most often temporal, but also frontal

DDx for seizure attacks

- ▣ TIA
- ▣ Syncope
- ▣ Migraine
- ▣ Movement disorders
- ▣ Panic attack
- ▣ Psychogenic seizure

Seizure vs syncope

Comparison of clinical features in cardiogenic syncope versus seizure disorders

Clinical features	Cardiogenic syncope	Seizure disorders
Loss of consciousness	Typical	Common
Episode duration	Seconds	Minutes
Involuntary movements	Common	Typical
Amnesia	Yes	Yes
Arrhythmia	Common	Rare*
Electroencephalogram	Slow waves Flattening	Focal or general spike activity
Responsive to AEDs	No	Often
Short term mortality†	High	Low

Seizure approach

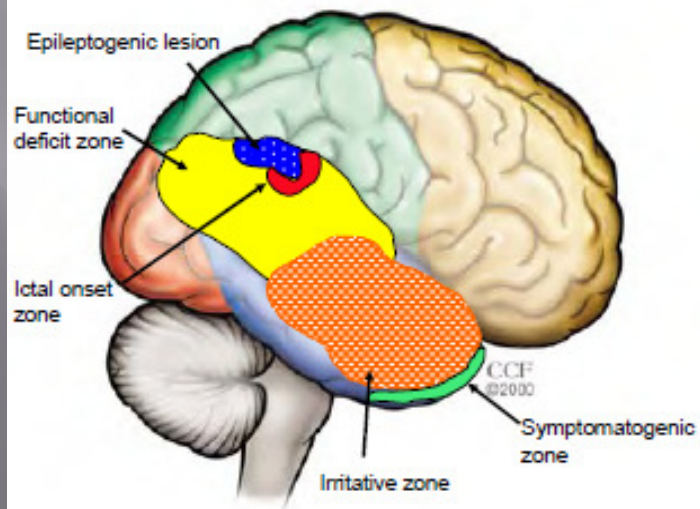
- ▣ **Non invasive tests**
 - Clinical history
 - MRI
 - video EEG
 - neuropsychological evaluation
 - nuclear medicine
- ▣ **Invasive monitoring**



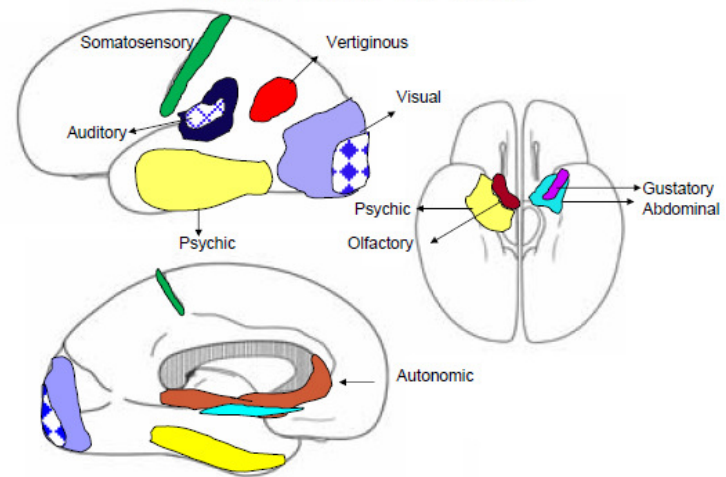
Clinical history



Defining the Epileptogenic Zone



Localization of Auras



Rona S. Textbook of Epilepsy Surgery, Lüders HO, ed. Informa Healthcare, 2008;432-42.

Questions that help clarify the type of seizure include the following:

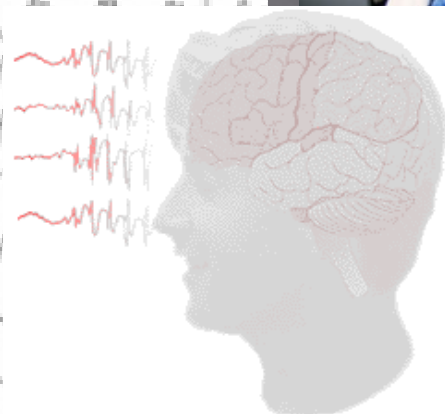
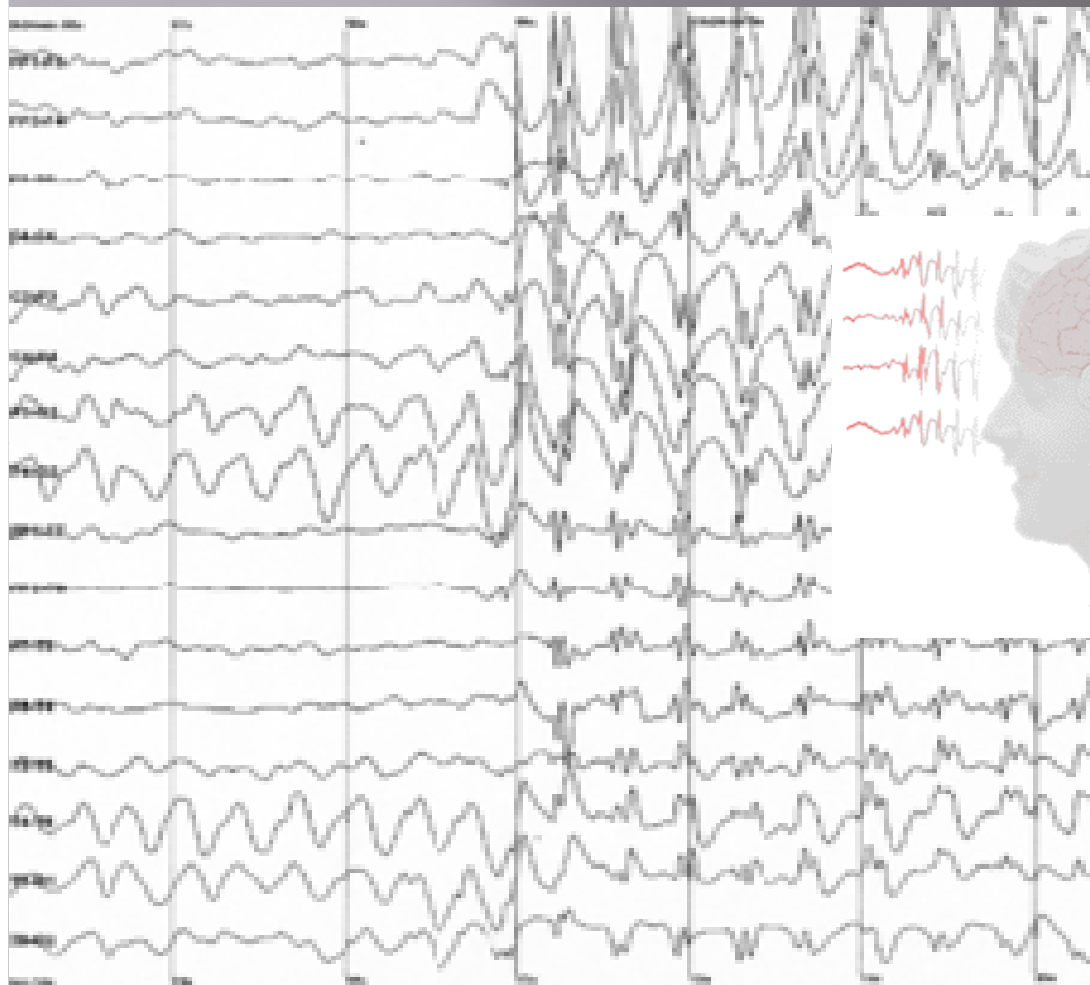
- ▣ Was any warning noted before the spell?
- ▣ What did the patient do during the spell?
- ▣ Was the patient able to relate to the environment during the spell ?
- ▣ How did the patient feel after the spell? How long did it take for the patient to get back to baseline condition?
- ▣ How long did the spell last?
- ▣ How frequent do the spells occur?
- ▣ Are any precipitants associated with the spells?

MRI

- Lesional
 - Tumor
 - Vascular
 - Trauma
 - Developmental
 - Mesial Temporal Sclerosis
- Non lesional



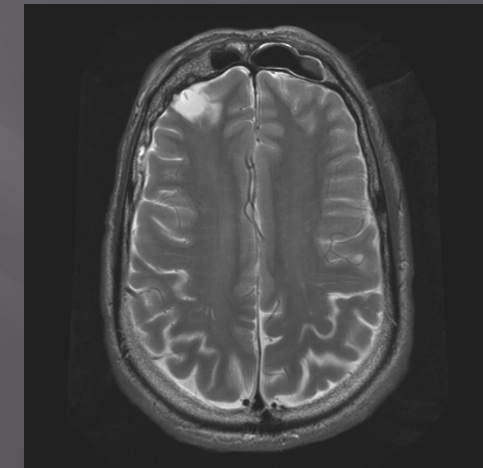
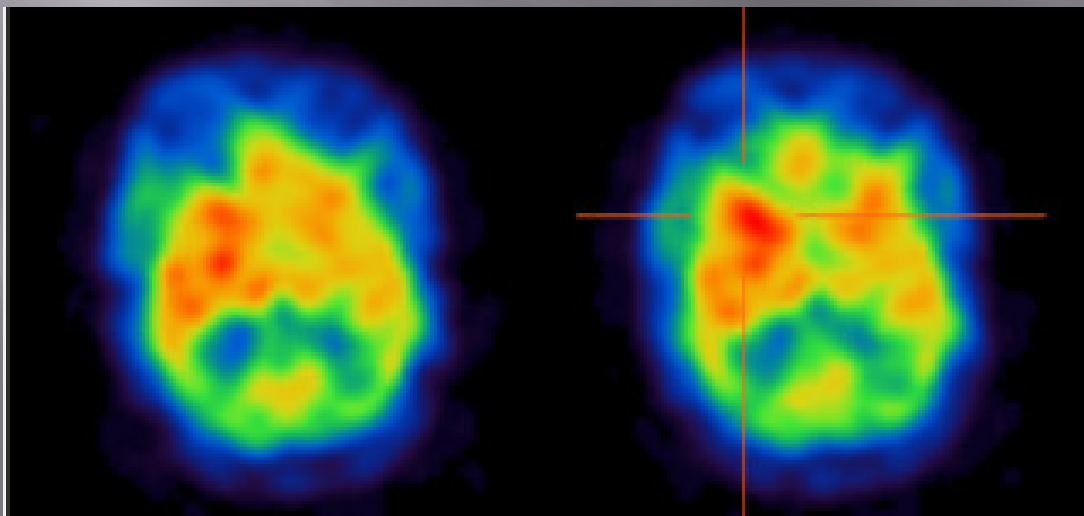
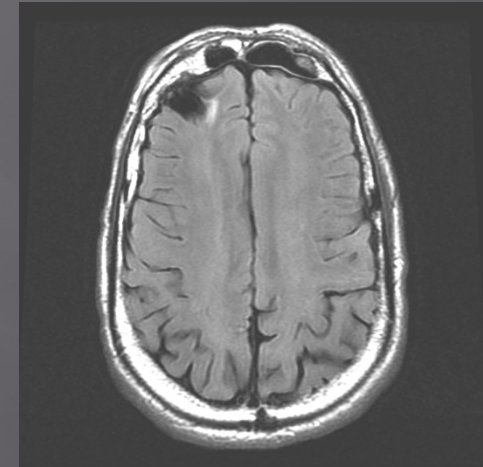
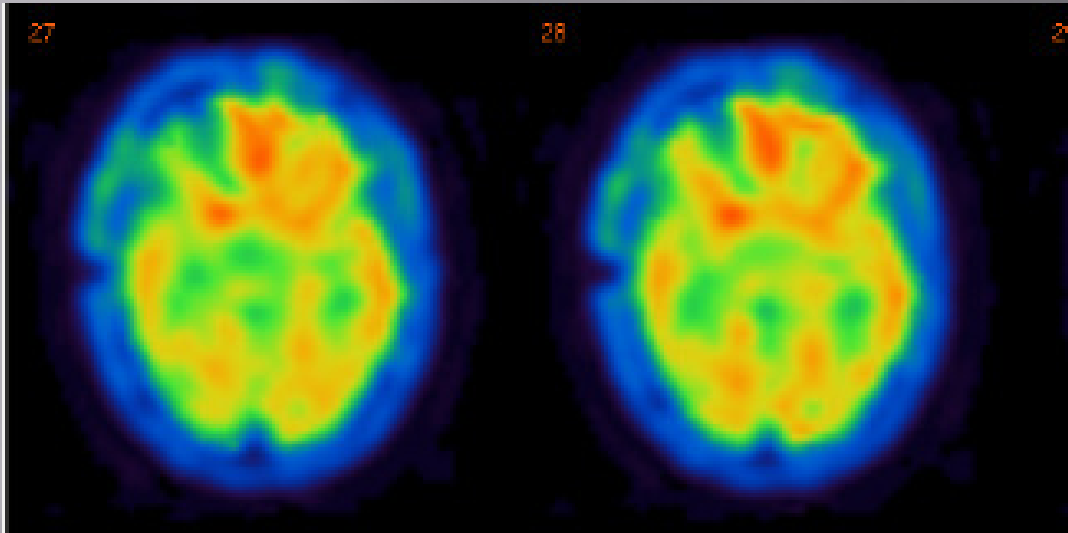
EEG



VEM



Nuclear Medicine



Cognitive Testing Neuropsychology

▣ Intelligence

▣ Memory

▪ Verbal

▪ Visual

▣ Language

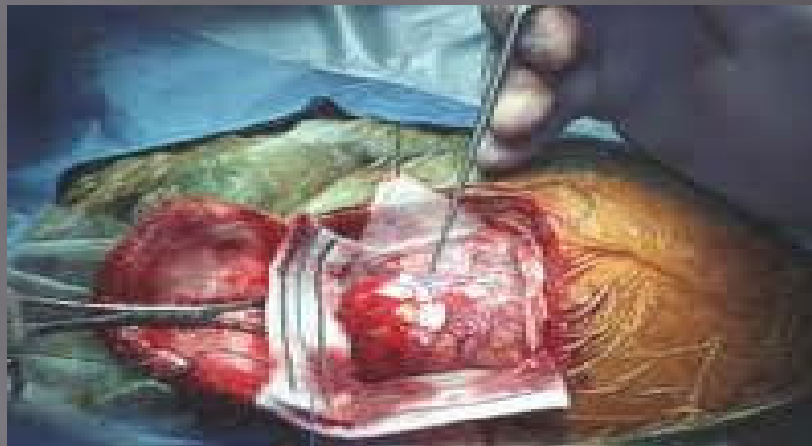


Treatment

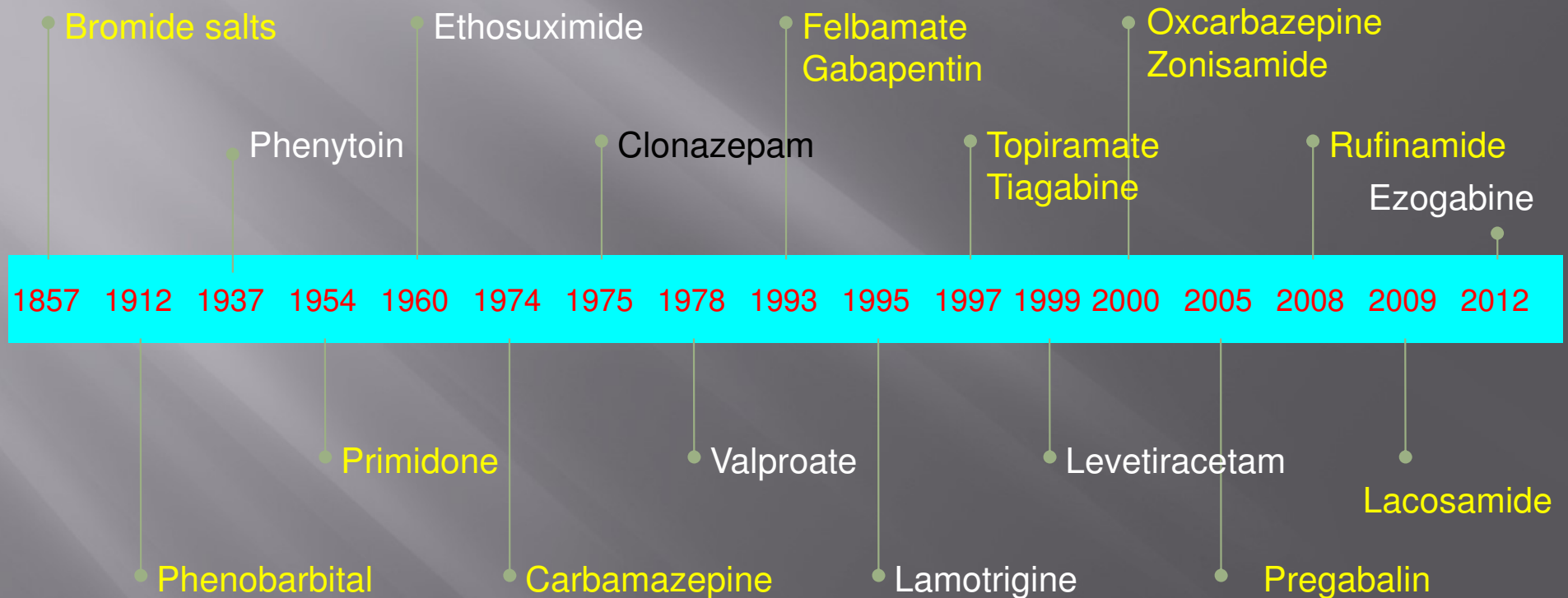
Medical



Surgical

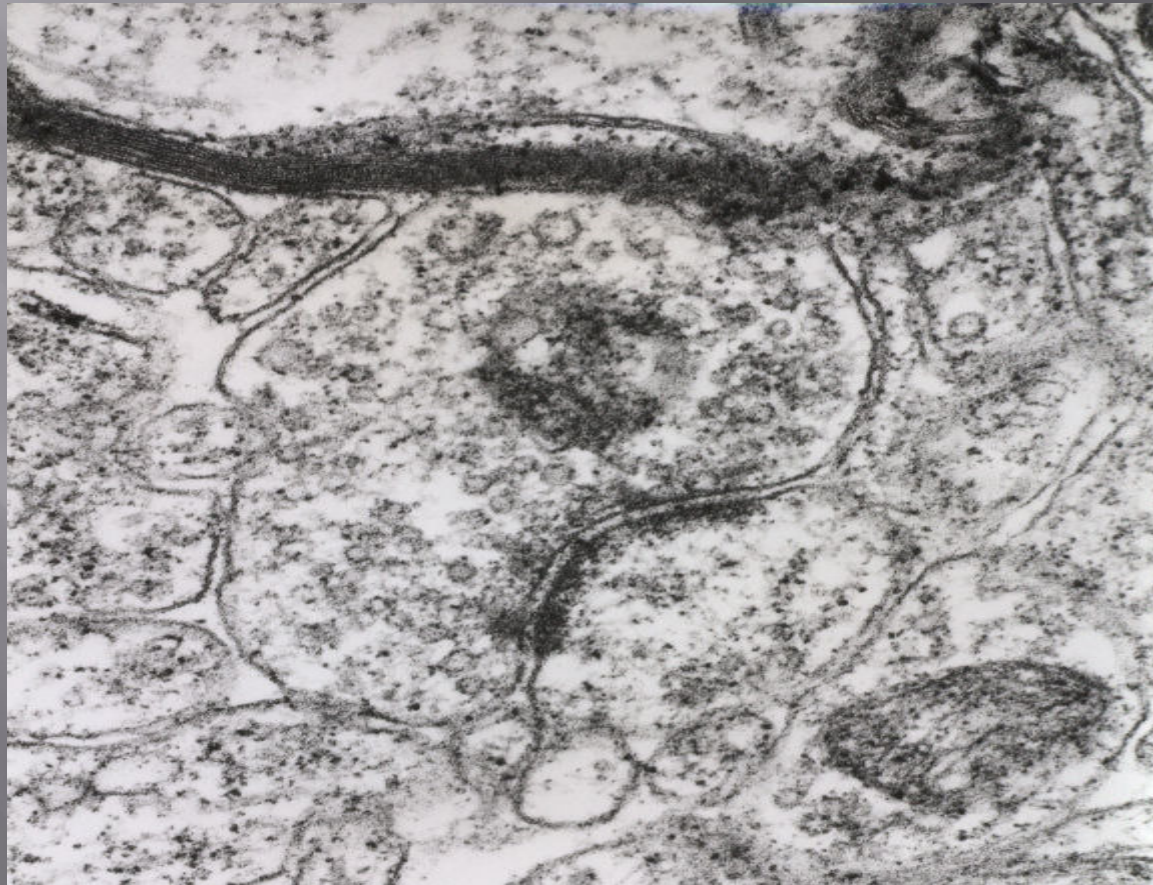


Epilepsy Drug Development



Mechanism of Action

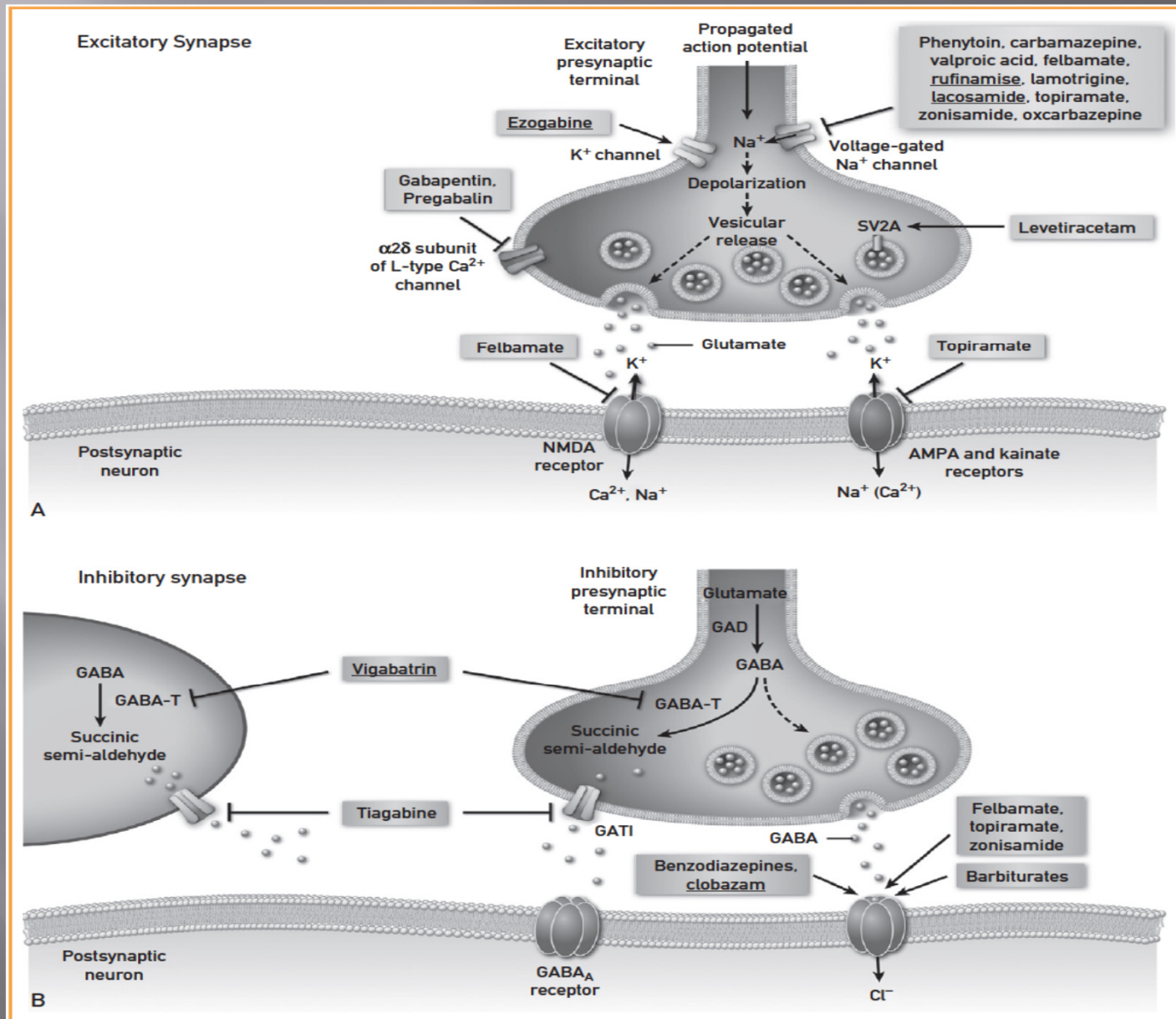
- *Current antiepileptic drugs are thought to act mainly by two main mechanisms:*



Mechanism of Action

- Reducing electrical excitability of cell membranes, possibly through inhibition of **sodium channel**.
- **Enhancing GABA**. This may be achieved by
 - inhibiting GABA-transaminase
 - Or
 - by drugs with direct GABA-agonist properties.

Mechanism of action of different AED



Clinical Uses of Antiepileptic Drugs

- ▣ *Tonic-clonic (grand mal) seizures*: phenytoin, valproate. Use of single drug is preferred when possible, because of risk of pharmacokinetic interactions.
- ▣ *Partial (focal) seizures*: carbamazepine, valproate; clonazepam or phenytoin are alternatives.
- ▣ *Absence seizures (petit mal)*: ethosuximide or valproate.
- ▣ *Myoclonic seizures*: valproate or clonazepam.

Basic rules for drug treatment

- ❑ Drug treatment should be simple, preferably using one anticonvulsant (monotherapy). “Start low, increase slow”.
- ❑ Add-on therapy is necessary in some patients...
- ❑ If patient is seizure-free for three years, withdrawal of pharmacotherapy should be considered.
- ❑ It should be performed very carefully and slowly! 20% of pts will suffer a further sz within 2 yrs.

Epilepsy treatment and pregnancy

- The risk of teratogenicity is well known (~5%), especially with valproates, but withdrawing drug therapy in pregnancy is more risky than continuation.
- **All** antiepileptic medications are not safe, however lamotrigine is the safest.
- Epileptic females must be aware of this problem and thorough family planning should be recommended.
- Over 90% of pregnant women with epilepsy will deliver a normal child.

Seizure Freedom with AED use

- ▣ 1st drug ----- seizure free (47%)
- ▣ 2nd drug----- seizure free (14%)
- ▣ 3rd drug----- seizure free (3%)

- ▣ Medication resistant 36%

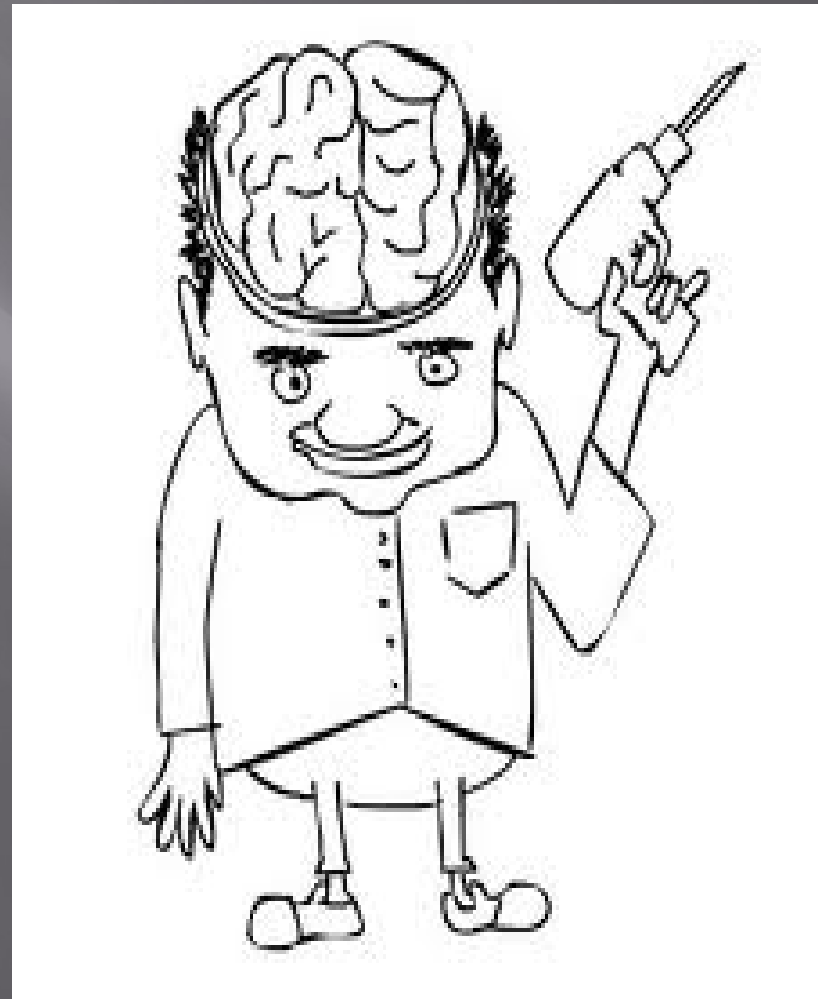
Drug resistant epilepsy

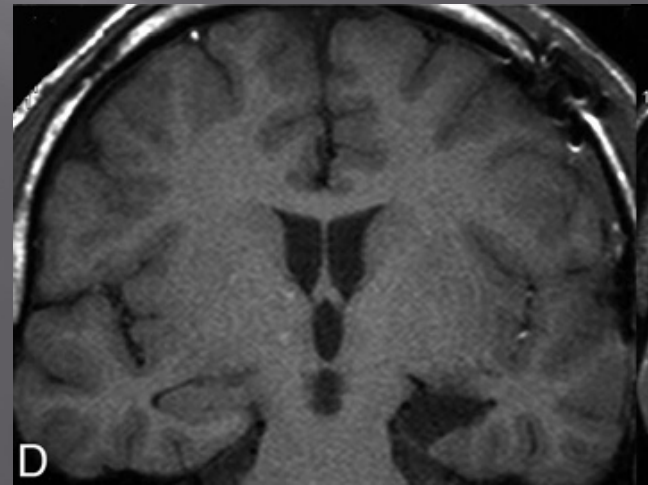
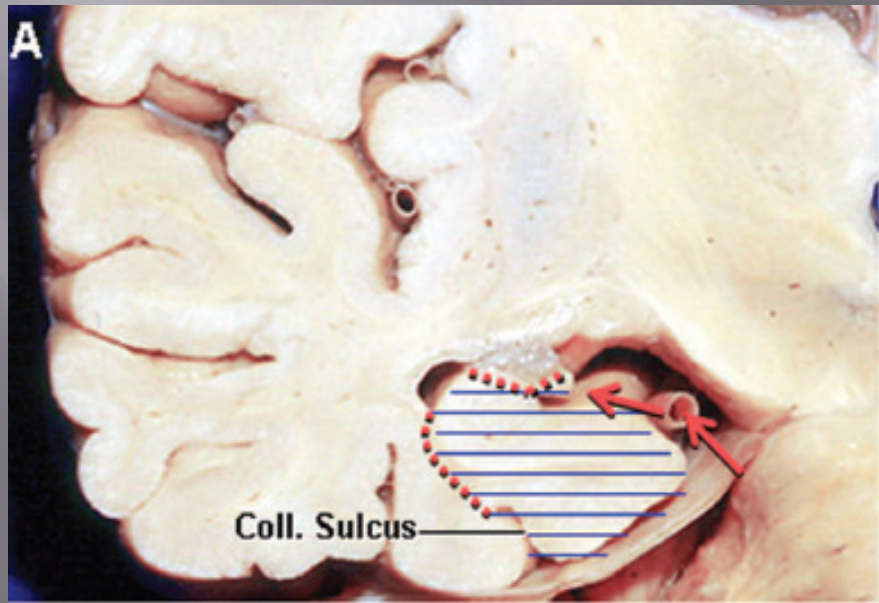
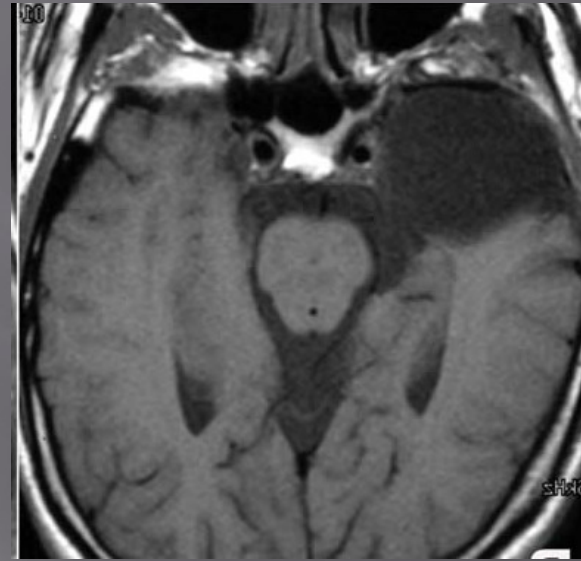
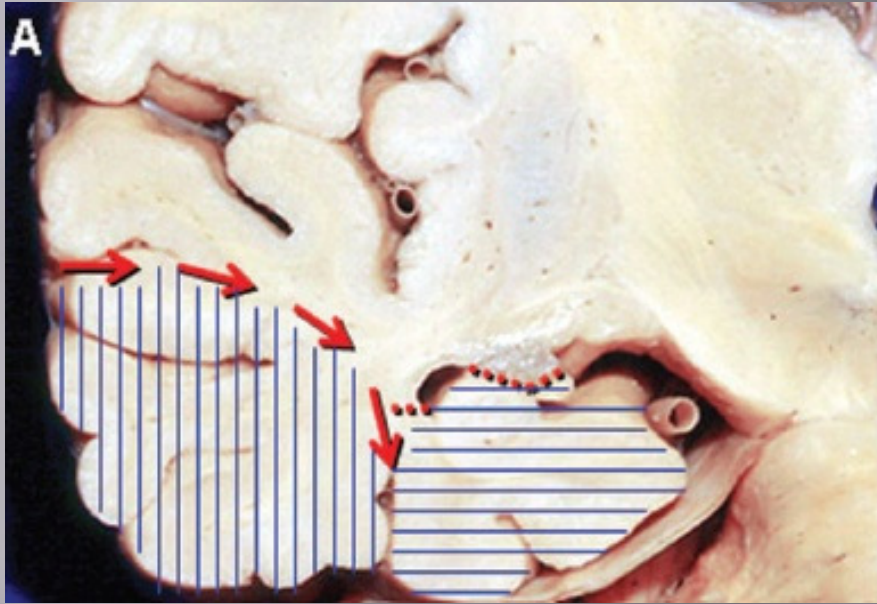
- Failure of at least TWO antiepileptic medications to completely control seizures
 - Appropriately chosen for seizure type
 - Taken as prescribed
 - Well tolerated (not failed due to side effects)

When to stop antiepileptic medications?

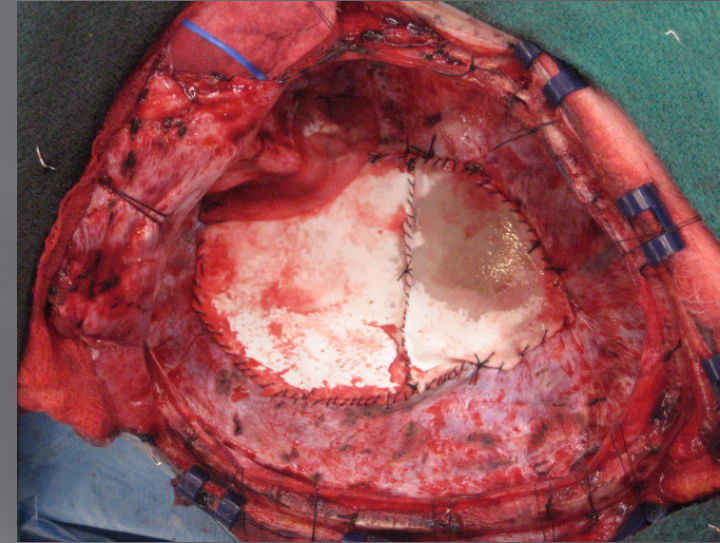
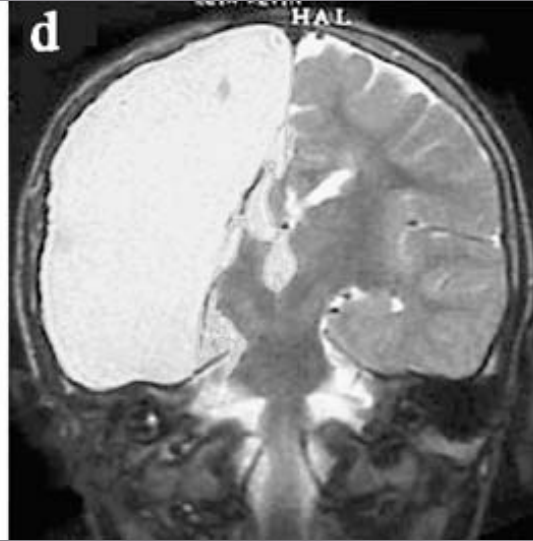
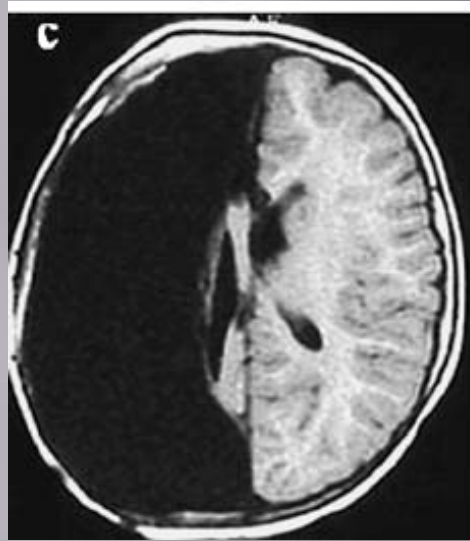


Surgery

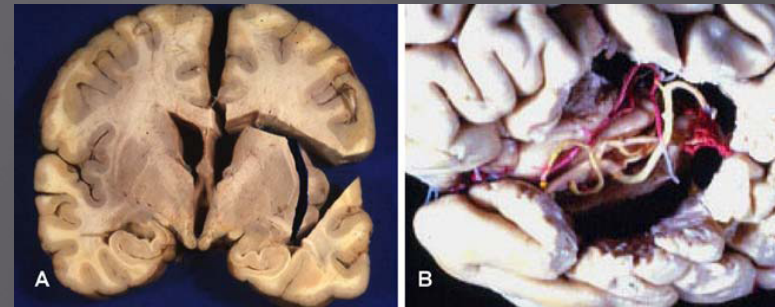
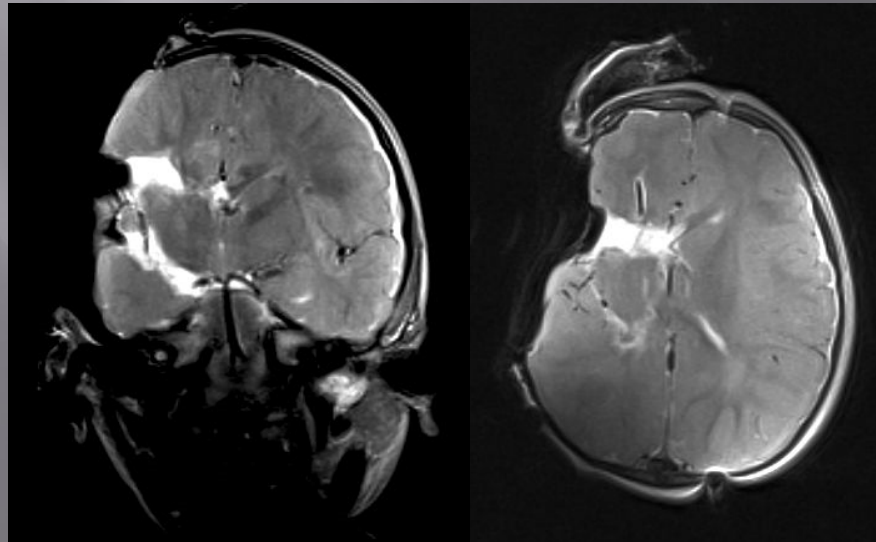




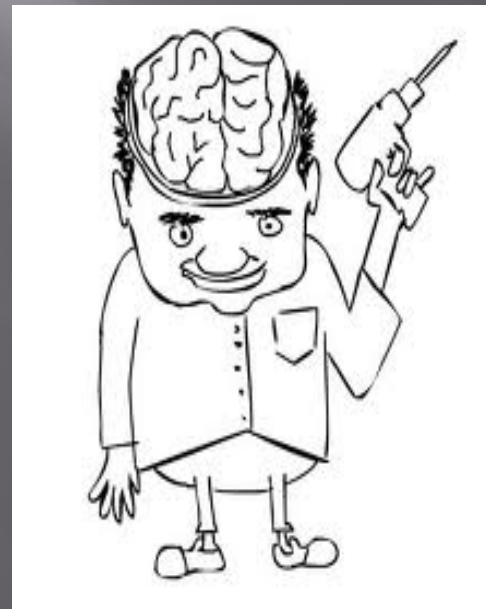
▣ Hemispherectomy



• Hemispherotomy



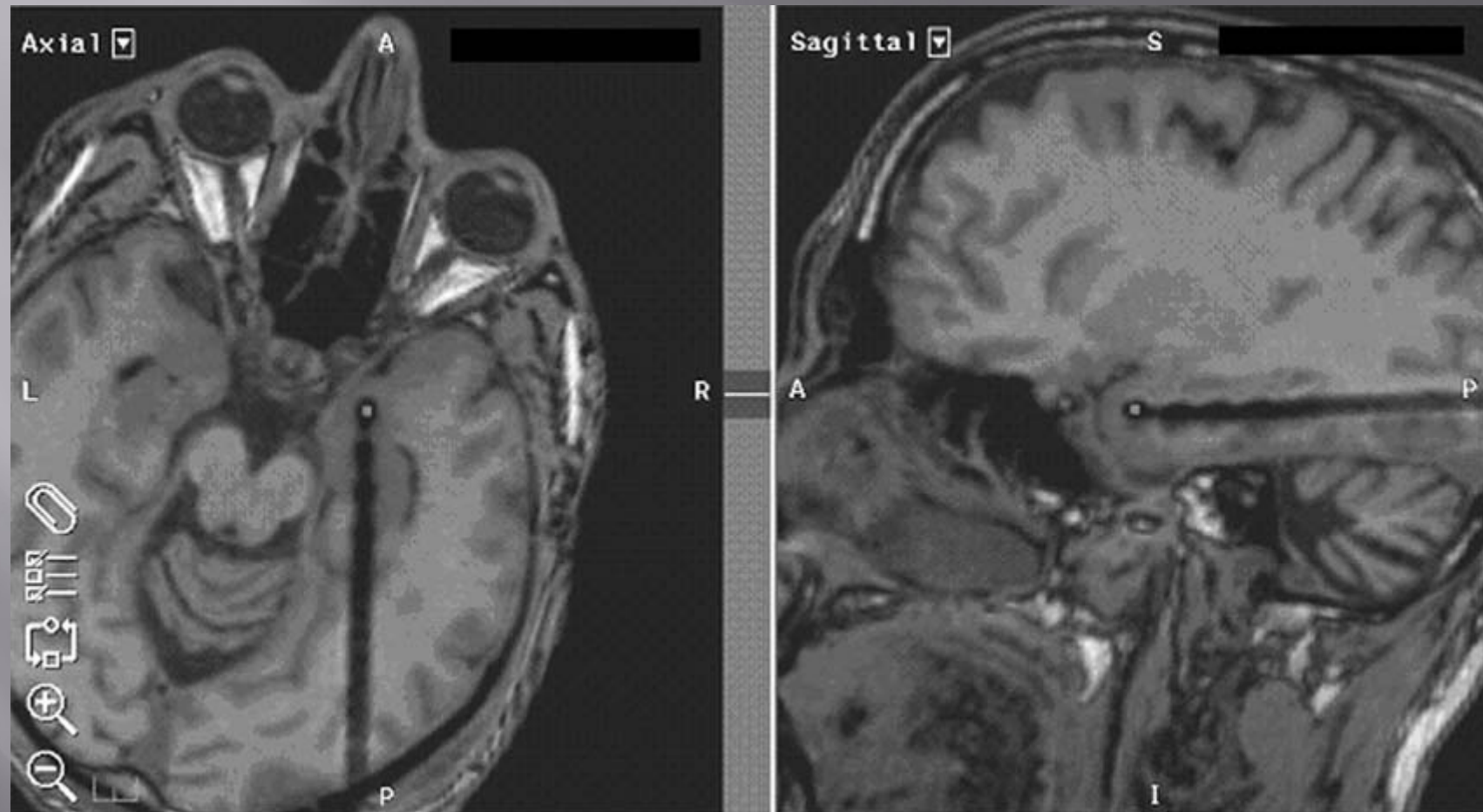
If my patient is not a good candidate
for surgery?



VNS



DBS





Thank
you

