Pathophysiology of Epilepsy



- * Seizures
- * Epilepsy

Definition

* Seizure (Convulsion)

- Clinical manifestation of synchronised electrical discharges of neurons
- * Seizures are symptoms of a disturbance in brain function, which can be due to epilepsy or other causes
- * A seizure is a sudden surge in electrical activity in the brain that causes an alteration in sensation, behavior, or consciousness

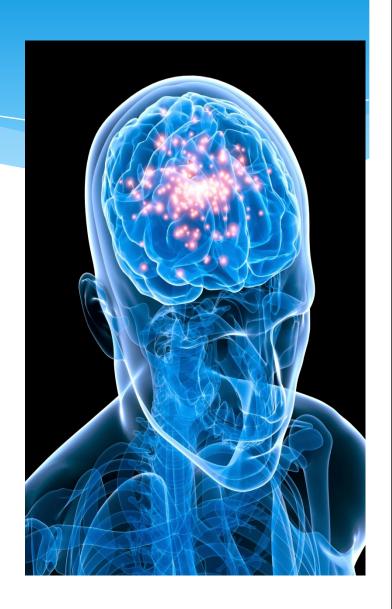
Epilepsy

Present when 2 or more unprovoked seizures occur at an interval greater than 24 hours apart

- * Sudden recurrent episodes of sensory disturbance
- * +/- Loss of consciousness, or convulsions
- * Associated with abnormal electrical activity in the brain
- * Abnormal, excessive electrical discharge of a group of neurons within the brain.
- * When a person has recurrent (2 or more), unprovoked seizures -> "epileptic".
- * Seizures is symptom of epilepsy

Provoked seizures

- Seizures induced by somatic disorders originating outside the brain
- * Fever
- * Infection
- * Syncope
- * Head trauma
- * Hypoxia
- * Toxins
- * Cardiac arrhythmias



Classification of Seizures

- * Seizures
- * Partial
- * or
- * Generalized

Partial (or Focal) Seizures

- Simple Partial
 - o Awareness <u>not</u> impaired
- Complex Partial
 - o Awareness impaired/lost
- Partial Seizures secondarily generalizing

Generalized Seizures

- Absence
 - o Typical
 - Atypical
- Myoclonic
- Clonic
- Tonic
- · Tonic-Clonic
- Atonic

* Focal / Partial seizures > their onset (start) is limited to part of the cerebral hemisphere

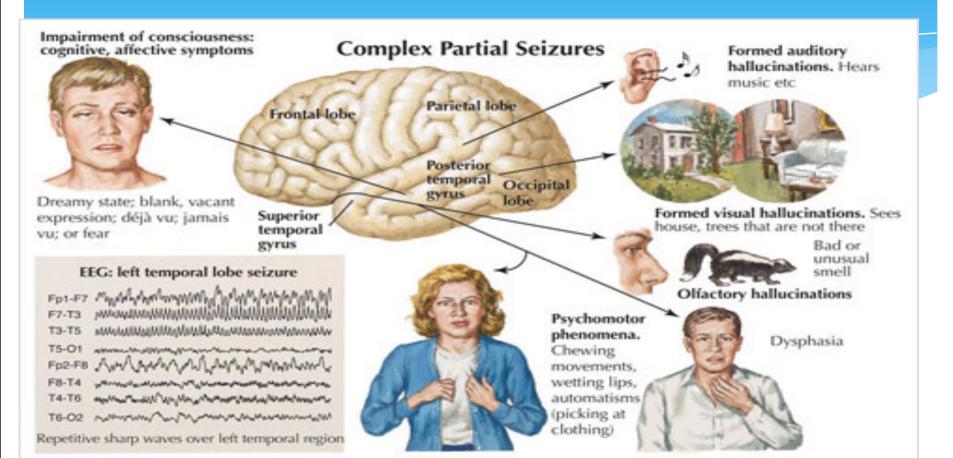
* Generalized seizures → those that involve the cerebral cortex diffusely (whole of it) from the beginning (generalized seizures)

partial seizures

<u>a. Simple partial seizures</u> manifest

motor, somatosensory, and psychomotor symptoms without impairment of consciousness

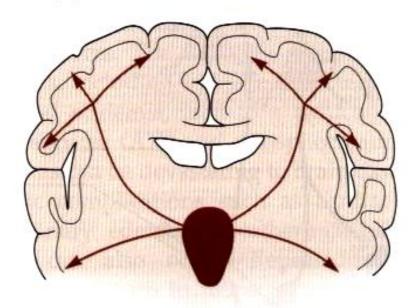
* b. Complex partial seizures manifest impairment of consciousness with or without simple partial symptoms

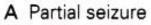


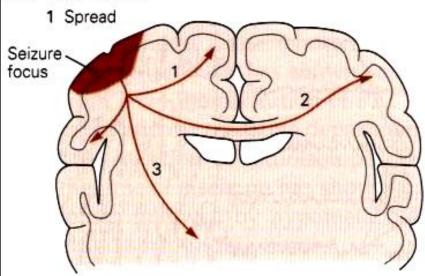
Generalized seizures

- * Manifest a loss of consciousness Concyulsive or non-convculsive
- * (1) Generalized tonic-clonic seizures (Grand Mal epileptic seizure)
- * (2) Absence seizures (Petit mal epileptic seizures)

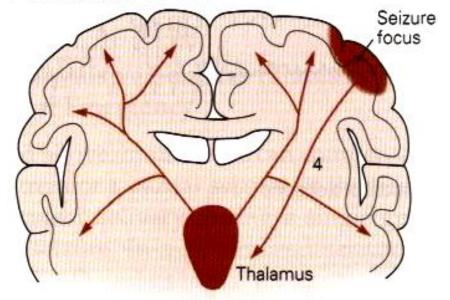
Primary generalized seizure







2 Secondary generalization

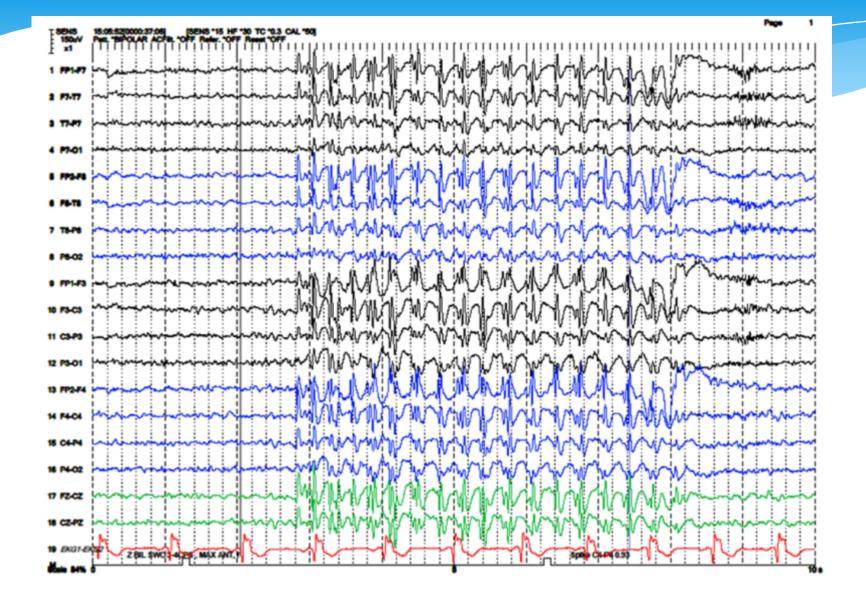


- * The onset of a seizures:
- Small group of abnormal neurons undergo
- Prolonged depolarizations
- Rapid firing of repeated action potentials
- * Spread to adjacent neurons or neurons with which they are connected into the process.

Generalized

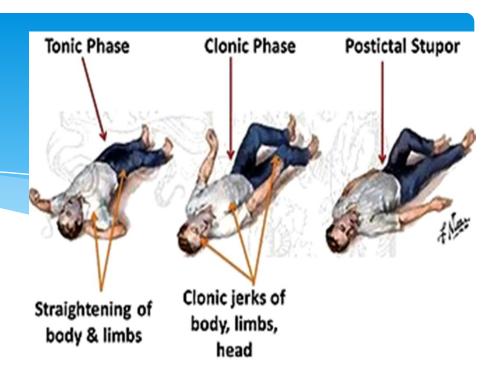
1- Generalized tonic-clonic (grand mal) seizure

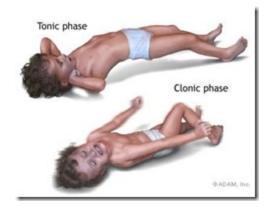
- * +/- aura
- * (peculiar sensation or dizziness; then sudden onset of seizure with loss of consciousness)
- * Tonic phase: Rigid muscle contraction in which clenched jaw and hands; eyes open with pupils dilated; lasts 30 to 60 seconds
- * Clonic phase: Rhythmic, jerky contraction and relaxation of all muscles in with incontinence and frothing at the lips; may bite tongue or cheek, lasts several minutes.
- * Postictal state: Sleeping or dazed for up to several hours.



A clinical seizure occurs
when the electrical
discharges of a large
number of cells become
abnormally linked together,
creating a storm of
electrical activity in the
brain.

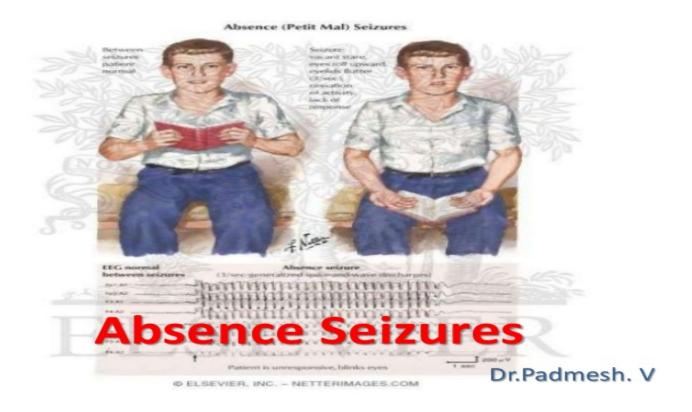
* Seizures may then spread to involve adjacent areas of the brain or through established anatomic pathways to other distant areas.





Absence (petit mal) seizure

- * a. Loss of contact with environment for 5 to 30 seconds.
- * b. Appears to be day dreaming or may roll eyes, nod head, move hands, or smack lips.
- * c. Resumes activity and is not aware of seizure.



* The <u>clinical manifestations of a seizure</u> reflect the area of the brain from which the seizure begins (i.e., seizure focus) and the spread of the electrical discharge.

- * Clinical manifestations accompanying a seizure are numerous and varied, including \rightarrow
- * (1) indescribable bodily sensations,
- * (2) "pins and needles" sensations,
- * (3) smells or sounds,
- * (4) fear or depression,
- * (5) hallucinations,
- * (6) momentary jerks or head nods,
- * (7) Déjà vu (over familiatry)
- * (8) staring with loss of awareness, and
- * (9) Convulsions → i.e., involuntary muscle contractions) lasting seconds to minutes.



Aetiology of seizures

- Epileptic
 - Idiopathic (70-80%)
 - Cerebral tumor
 - Neurodegenerative disorders
 - Secondary to
 - Cerebral damage: e.g. congenital infections, intraventricular hemorrhage
 - Cerebral dysgenesis/malformation: e.g. hydrocephalus

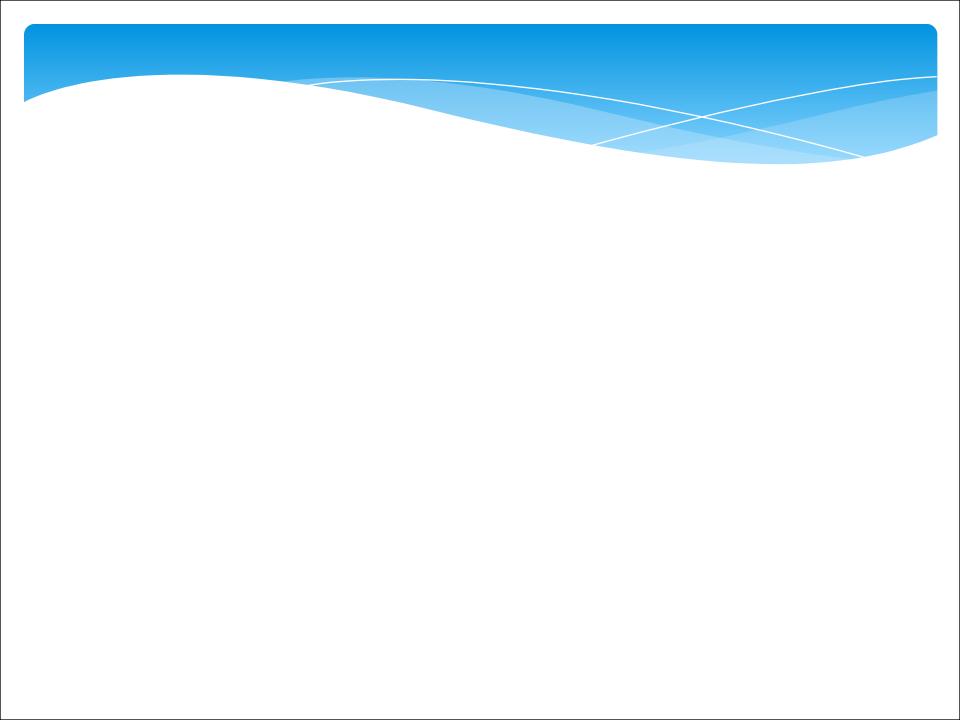
Aetiology of seizures

- Non-epileptic
 - Febrile convulsions
 - Metabolic
 - Hypoglycemia
 - HypoCa, HypoMg, HyperNa, HypoNa
 - Head trauma
 - Meningitis
 - Encephalitis
 - Poisons/toxins

Pathophysiology of Epilepsy (at molecular level)

- Cortical cell membrane level
- ➤ Instability of the nerve cell membrane → Polarization abnormalities (excessive polarization ,hypopolarization , or lapses in repolarization), allowing the cell to be more susceptible to activation → Hypersensitive neurons with lowered thresholds for firing and firing excessively , related to →

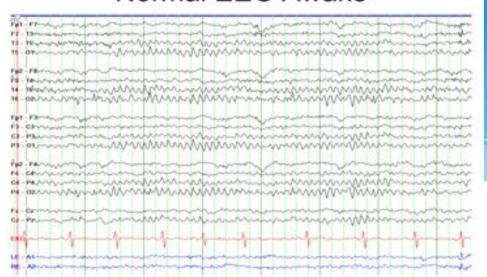
- (1) Excess of Excaitatory (acetylecholine- or Glutamate related activity)
- (2) Decreased inhibitory (GABA -related activity)
- ➤ Together and/or (2) above → leading to instability of cell-membrane & lowered threshold for exciatation → excessive polarization, hypopolarization allowing the cell to be more susceptible to activation spontaneously or by any ionic imbalances in the immediate chemical environment of neurons



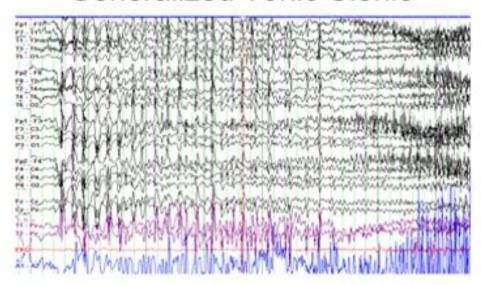
* Electroencephalogram (EEG)

- * EEG >>>> diagnosis, classifying seizures >>>> therapeutic decisions
- * spikes or sharp waves (Epileptiform EEG patterns)
- > Focal epileptiform discharges indicate focal epilepsy
- Generalized epileptiform activity indicates a generalized form of epilepsy.

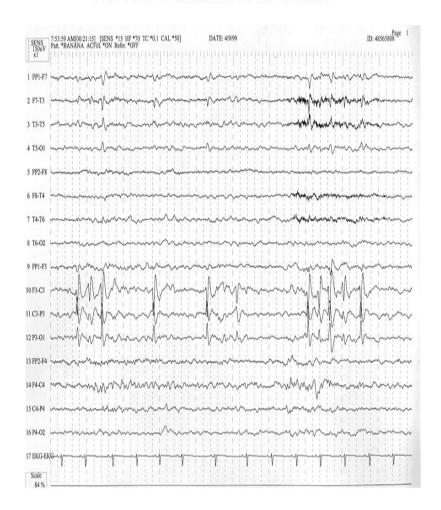
Normal EEG Awake



Generalized Tonic Clonic



EEG: Partial Seizures



- * Some types linked to genes
- * (run in families)
- Genetic abnormalities >>>> increasing a person's susceptibility to seizures_that are triggered by an environmental factor
- * Several types of epilepsy have now been linked to defective genes for ion channels

Pathophysiology

- * Genetic factors
 - * At least 20 %
 - * Some examples
 - * Benign neonatal convulsions -- 20q and 8q
 - * Juvenile myoclonic epilepsy--6p
 - * Progressive myoclonic epilepsy--21q22.3

