

Neonatal Seizures



KEMRI Wellcome Trust



Objectives

• Review the properties of commonly available

anticonvulsants; Phenobarbitone, Levetiracetam & Phenytoin

• Illustrate the need for appropriate supportive care

during a convulsion



Introduction

- About 4 out 10 neonates with seizures progress to status epilepticus.
- Timely and correct treatment of seizure is desired
- Higher seizure burden is associated with worse outcome.
- Seizures become more difficult to treat the longer they last.
- Only half of neonates treated with current single anticonvulsant drugs respond to treatment while another 10-20% respond with multiple drugs.



Introduction

Causes of	Neonatal Seizures
Hypoxic ischemic encephalopathy	Commonest cause in above 1500grams
Intraventricular Hemorrhage	Commonest cause below 1500gms
Stroke	Arterial or venous
Acute Metabolic disorders	Hypoglycemia, hypocalcemia, hypomagnesemia
CNS Infections	Meningitis, encephalitis, intrauterine infections
Congenital Brain Malformations	Lissencephaly,holoproncephaly

Definitions

Clonic Seizures

 Rhythmic jerking, consciousness usually preserved.
Focal, multifocal or generalised. Often correlate with a structural lesion.

Tonic Seizures

 Extension of upper and lower limbs accompanied by pronation of arms and clenching fists. Often less than a minute and seen most commonly in the first 24 hrs of life following an hypoxic event



Definitions

Subtle seizures (motor automatism & autonomic signs)

- Motor automatisms (such as chewing, swallowing, sucking, repetitive tongue movements, "cycling", "boxing", "pedaling", "swimming" Eyelid fluttering, eye deviation, fixed open stare, chewing, sucking, tongue thrusting,)
- Autonomic signs (changes in heart rate or breathing pattern, flushing, salivation, pupil dilatation, tachycardia, BP instability and apnoea)



Principles of Managing Seizures

- Resuscitation and supportive measures
- Assessment and treatment of the underlying cause
- Anticonvulsant drugs
- Wean of anticonvulsant drugs
- Follow-up and rehabilitation



Managing the risks of seizures and their treatment

Airway

- Positioning
- Suction
- Support after seizure

Breathing

- Start on Oxygen
- Check after seizure

Circulation

- Temperature gradient?
- Severe Pallor?

Disability

- What drugs have been used?
- Glucose?
- Calcium
- Need of phenobarbitone



The commonly used drugs cause respiratory depression Anticipate – be prepared to intervene



Phenobarbitone

- First line treatment for neonatal seizures
- It is cheaper and easily available that phenytoin
- Time period of phenobarbital distribution is estimated 30minute
- Additional seizure drug doses should <u>not be given sooner</u> <u>than 60 minutes from administration of the IM injection to</u> allow for evaluation of the full medication effect prior to administering further doses.
- Phenobarbitone is easier to administer with one daily dose being adequate after attainment of therapeutic level.



Phenobarbitone – side effects

- Respiratory depression
- CNS depression
- In overdose or rapid iv infusion coma and hypotension.

Monitor respirations, pulse and blood pressure

PS: In the absence of clinical seizures, neonates with hypoxic - ischemic encephalopathy need not to be given prophylactic treatment with phenobarbital



Phenytoin

- Phenytoin and phenobarbitone are equally effective in neonatal seizures
- Has unpredictable pharmacokinetics and narrow therapeutic index. The level between efficacy and toxicity is very narrow and thus requires very close blood level monitoring
- Has more severe side effects; cardiac arrythmia requires cardiac monitoring during administration, extravasation



Phenytoin

- Not ideal medication for maintenance on discharge; Has erratic oral absorption and variable metabolism with age of the young infants
- Phenytoin only provide 10-15% of increase in seizure control when given following phenobarbitone failure



Levetiracetam

- Inhibits burst firing without affecting normal neuronal excitability
- May selectively prevent hypersynchronization of epileptiform burst firing and propagation of seizure activity.
- Rapidly and completely absorbed even after oral administration (peaks1-1.5hrs, duration 12hrs)
- Side effects;
 - Neuropychiatric symptoms, weakness, drowsiness, dyskinesia, fatigue.
 - Stevens-johnson syndrome, toxic epidermal necrolysis.
- Dilution: Dilute 30mg/kg in 10 mL of 0.9% NaCl, D5W, or RL.
- Infuse over 15 min



Managing Seizures





Managing Seizures – if no response to phenobarbitone



In the absence of clinical seizures, neonates with hypoxic-ischaemic encephalopathy need not to be given prophylactic treatment with phenobarbitone



When to stop anticonvulsants

- Stop anticonvulsants if neonate has/is;
 - Normal neurological examination
 - Normal electroencephalography
 - Seizure-free for >72 hours
- Restart the drug(s) in case of recurrence of seizures
- In neonates in whom seizure control is achieved with a single anticonvulsant drug; discontinue drug abruptly without any tapering of the doses.



When to stop anticonvulsants

- In neonates requiring more than one anticonvulsant drug for seizure control;
 - Stop drugs one by one, with phenobarbital being the last drug to be withdrawn.
- Neonates with abnormal neurological examination;
 - Continue phenobarbitone for ONE months then review





Questions





Summary

- Neonatal seizures are symptoms of underlying CNS/systemic condition – assess the patient! – remember hypoglycaemia
- Phenobarbitone is first line treatment and levetircateam second line.
- 3. Stop anticonvulsant if neonate seizure free for 72hrs

