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ENLS Version 5.0 Status Epilepticus

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Editors' Note: Global Considerations

The intent of the editors, authors, and reviewers of this ENLS topic was not to address all the variations in international practice for the different diseases. We have discussed major practice variances (e.g., the availability of diagnostic testing, or the type of medications used) and encourage learners to use the ENLS algorithms as a framework on which any relevant local practice guidelines can be incorporated.







Status Epilepticus

Learning Objectives

- Diagnose status epilepticus
- Implement Emergent Initial Therapy
- Implement Urgent Control Therapy
 - Identify and treat Refractory Status Epilepticus



Diagnosis of Status Epilepticus

Definition: 5 minutes or more of:

- Continuous seizure activity, OR
- Recurrent seizure activity without return to baseline.





Etiologies of SE

Chronic Processes

Metabolic disturbances: electrolyte abnormality, renal failure

Sepsis

CNS infection: meningitis, encephalitis, abscess

Stroke: ischemic, ICH, SAH

Acute Processes

TBI

Drugs (medication NON-ADHERENCE, intoxication, or withdrawal from benzodiazepines, barbiturates, ETOH, and other sedatives)

Cardiac arrest, hypoxia

Hypertensive encephalopathy, PRES

Autoimmune encephalitis, paraneoplastic



Remote CNS pathology (e.g. stroke, abscess)

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Checklist for the first hour

Checklist

- □ Fingerstick Glucose
- □ Obtain IV Access
- □ Pulse Oximetry, BP and cardiac monitoring, Supplemental O2 and Fluid as needed
- □ Labs: CBC, BMP, Ca, Mg, HCG in females of childbearing age
- \Box Possible CT head
- □ Continuous EEG (if available); notify EEG tech if available (as soon as available unless patient returns to pre-status epilepticus baseline)
- □ Consider rapid-response EEG with limited montage if continuous EEG is not available





Initial work-up in ER

All patients

- Airway: adjuncts, O₂
- Monitor EKG, BP, O₂
- IV or IO access: IVF
- Treat hypoglycemia
- Labs: blood glucose, CBC, BMP, Ca, Mg

Consider





LP: if any suspicion for infection





Treat Fast



Lowenstein Neurology 1993; 43:483-488.



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- A 28-year-old man with no known past medical history is brought by EMS to the emergency department after suffering a seizure at work.
- EMS did not administer any medication in the field or en route to the hospital.
- Seizure activity started approximately 22 minutes ago and continues.

What is the appropriate first line pharmacologic agent to treat this patient's status epilepticus?





- A. Levetiracetam 1000 mg IV over 15 minutes
- B. Lorazepam 2 mg IVP
- C. Lorazepam 0.1 mg/kg IVP
- D. Propofol 1 mg/kg IV push followed by infusion of 0.5 mg/kg/hr
- E. Phenytoin 20 mg/kg IV push





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The Veterans administration (VA) cooperative trial identified 0.1 mg/kg of IV lorazepam as the preferred first line pharmacologic agent.

IM midazolam may also be considered as a first line pharmacologic agent.







Silbergleit et al, NEJM 2012



Emergent Initial Therapy

- Seizure activity needs to be treated as fast as possible
- Benzodiazepines are first line therapy
 - IV access will determine the benzo of choice
- Establish IV access as soon as possible



Emergent Initial Therapy

Recommended dosing

NO IV ACCESS

• Midazolam 10mg IM if > 40kg

Adults • Midazolam 5mg IM if < 40kg

• If unavailable, Diazepam 20mg PR

- Midazolam Buccal 0.2-0.5 mg/kg (max 10mg)
- Midazolam Intranasal 0.2mg/kg (max 10mg)
- Midazolam IM 0.2mg/kg
 - >40kg: max dose 10mg
 - 13-40kg: max dose 5mg





Children

Emergent Initial Therapy

Recommended dosing

WITH IV ACCESS

All Patients • Lorazepam 0.1mg/kg (up to 4mg per dose)





50-year-old man presents by EMS for seizures lasting 14 minutes which stopped for 2 minutes following lorazepam 4 mg x 1 dose.

The seizures then recurred without return to baseline.

Does this patient meet criteria for the diagnosis of status epilepticus?



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- A. No, 30 minutes of continuous seizure activity is required before a patient is considered to be in status epilepticus
- B. No, the seizure stopped for a short period of time indicating recurrent seizures but not status epilepticus
- C. No, the diagnosis of status epilepticus cannot be reached without confirmatory EEG testing
- D. Yes, the patient's seizure activity lasted longer than 5 minutes without return to neurologic baseline
- E. Yes, the patient has a history of seizure activity and is therefore in status epilepticus





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- B. No, the seizure stopped for a short period of time indicating recurrent seizures but not status epilepticus
- C. No, the diagnosis of status epilepticus cannot be reached without confirmatory EEG testing
- D. Yes, the patient's seizure activity lasted longer than 5 minutes without return to neurologic baseline
- E. Yes, the patient has a history of seizure activity and is therefore in status epilepticus







Urgent Control Therapy

- If SE persists after 10-20 mins despite two adequate doses of benzos, second-line antiseizure medications are added
- If SE is aborted, antiseizure medications are added to prevent further seizures



Urgent Control Therapy

Adults and Children

Fosphenytoin IV 20mg PE/kg

Valproate IV 40mg/kg

Levetiracetam IV 60mg/kg (max dose 4500mg)

Neonates

Phenobarbital IV 20mg/kg

Fosphenytoin IV 20mg/kg

Levetiracetam IV 40-60mg/kg (max dose 4500mg)





Have Seizures Stopped?





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Case 2, continued

- The patient continues to have convulsions intermittently
- Not following commands in between seizures
- The patient is now in refractory SE



Treatment of Refractory SE

Adults

Midazolam	 Bolus: 0.2 mg/kg IV Infusion: 0.1-2.0 mg/kg/h
Propofol	 Bolus 1-2 mg/kg IV Infusion: 20-200 mcg/kg/min
Ketamine	 Bolus: 1-2 mg/kg IV Infusion: 0.5-10 mg/kg/h



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Treatment of Refractory SE

Children

Midazolam	 Bolus: 0.1-0.2 mg/kg IV Infusion 0.1-1.0 mg/kg/h
Phenobarbital	 Bolus: 20 mg/kg IV Infusion: n/a
Ketamine	 Bolus: 1-2 mg/kg IV Infusion: 0.5-10 mg/kg/h



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 He is intubated using propofol and rocuronium

 No more convulsions are seen

• EEG is started





EEG monitoring

- EEG is necessary to evaluate for non-convulsive SE in patients not returning to baseline
 - Ideally, EEG would be initiated <u>within one hour</u> of suspected nonconvulsive SE or if the patient does not return to baseline
 - Rapid response EEG with reduced montage may aid in quickly identifying patients with non-convulsive SE in critical care settings where continuous EEG monitoring is not available
- cEEG is necessary to assess treatment goals of sedatives in the management of refractory SE
 - Titration goal controversial seizure control, burst suppression, or completely suppressed background





Pediatric Considerations

- Similar etiologies as adults, but also include
 - Febrile seizures (30-50% pediatric SE)
 - Genetic and metabolic disorders
 - Hyponatremic seizures
 - Hypoparathyroidism and hypocalcemia (first weeks of life)
 - Abuse head trauma
- Neonatal seizures are often non-convulsive





Pediatric Considerations

• First-line emergent treatment similar to adults

- Refractory treatment
 - Favor midazolam and pentobarbital over propofol
 » Higher risk of propofol infusion syndrome





Nursing Considerations

- Knowledge of action, timing and side effects of antiseizure medications
- Safety measures
 - Lateral recumbent position
 - Head of bed elevated
 - Padding
 - Suction supplies
 - No restraints or putting anything into the mouth
 - Supplemental O2 if hypoxic
 - Vigilant monitoring of vital signs
- Obtain IV access





Pregnancy Considerations

- Treatment similar, with Benzo as first line
 - Exception is Eclampsia give magnesium
- Caution with the teratogencity of antiseizure medications
 - High Risk = Valproate, phenobarbital, and phenytoin
 - Lower Risk = lamotrigine, levetiracetam
- Monitor drug levels



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Communication

- □ Clinical presentation
- □ Duration of status epilepticus
- □ Relevant PMH/PSH
- Relevant labs, including anticonvulsant levels if drawn
- Prior medications, medication given so far
- □ Neurological examination
- \Box Brain imaging/LP results (if available)

Example of a sign-off narrative:

- 54 yo M found on sidewalk, shaking with all 4 extremities, no commands, Lac R forehead.
- Unclear duration of SE but seizing on off for 25 min
- Hx of epilepsy after remote TBI, with subtherapeutic levels of phenytoin
- Received lorazepam 4 mg IV x 2 and loaded with 20 mg/kg fosphenytoin
- Vital signs stable, afebrile
- Exam: not shaking but comatose, treated with propofol and rocuronium for intubation
 - Head CT no evidence for bleed, stroke, or mass

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Clinical Pearls - Management

- Early, aggressive management of patients with SE is imperative. Delay of therapy reduces the likelihood of seizure termination.
- Benzodiazepines are frequently underdosed for initial therapy, and should be dosed aggressively in SE adequately
- Continuous EEG is necessary
- Diagnostic work-up in parallel to treatment





Clinical Pearls – Medication

- Benzo's are first line treatment
- Fosphenytoin is the prodrug of phenytoin; it can be administered more rapidly than phenytoin
- Fosphenytoin, levetiracetam, and valproic acid have comparable rates of seizure cessation in patients with SE who have failed a benzodiazepine (<50%).
- Monitor antiseizure medication levels and interactions with other mediccations and diet





Questions?

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