Clinical Policy: Critical Issues in Evaluation and Management of Adult Patients Presenting to the ED with Seizures (Non-Status Epilepticus)

Written By: Sarah Schuman Harlan, PharmD; Edited By: Jessica Winter, PharmD, BCPS, BCCCP

In patients with known seizure disorders, it is appropriate to resume either IV or PO route of administration of the home ASD *(Level C Recommendation)*. There is no recommendation that loading ASDs after noncompliant seizure (non-status epilepticus) is necessary, however available loading doses are supplied below if loading is clinically appropriate. An alternative option includes resuming home maintenance dose as prescribed.

Drug	Non-Status Epilepticus Loading Dose	Status Epilepticus Loading Dose*	Adverse Events	Seizure Recurrence rate (non-status epilepticus)	Notes
Carbamazepine Tegretol, Equetro	8 mg/kg PO (single oral load) using oral suspension; max 1600 mg ¹	Refractory SE: 8 mg/kg PO (use oral suspension); max 1600 mg	58% experienced: drowsiness (26%), nausea (23%), dizziness	Not Studied 93% achievement of therapeutic conc	PO tablet has slow/erratic absorption
Gabapentin Neurontin, Gralise	900 mg/d PO (300 mg TID) x 3 days ²	Refractory SE: No load; start at 300 mg Q 8 hours	Somnolence, dizziness, ataxia and fatigue	No difference from slower load (only assessed 5 day period)	Adjunct for partial seizures
Lacosamide Vimpat	Loading dosage not studied; Maximum oral absorption 800 mg ³	Refractory SE: 5-10 mg/kg IV over 5-10 min (max 500 mg)	Mild-mod dizziness, HA, back pain, somnolence, injection site pain	Not Studied	Adjunct for partial seizures; withdrawal seizures with abrupt discontinuation
Lamotrigine Lamictal	6.5 mg/kg single oral load in on LMG for >6 months w/o hx of intolerance and only off LMG <5 days ⁴	Not recommended	Mild, transient nausea	Not Studied	Frequent and serious rashes; DO NOT load if hx of rash or patient not previously on LMG
Levetiracetam Keppra	1500 mg PO load Rapid IV loading up to 60 mg/kg well tolerated ^{5,6}	First Line: IV: 60 mg/kg (over 10 min); Max 4500 mg	Fatigue, dizziness, rare pain at injection site	No seizures within 24 hours of loading in study of oral load	
Phenytoin Dilantin, Phenytek	20 mg/kg divided in max 400 mg PO q 2 hours or 18 mg/kg IV (max rate 50 mg/min) ^{7,8,9}	Not recommended	IV faster to load but more serious ADE (low BP, bradyarrhythmias, cardiac arrest, extravasation)	No significant difference between PO and IV loading	Oral cheaper but takes >5 h to reach therapeutic levels; IV requires filter, infusion pump
Fosphenytoin Cerebyx	18 PE/kg max rate 150 PE/min ^{7,9}	First Line: IV: 20 PE/kg (rate up to 150 mg/min); Max 2000 mg	Fewer ADE compared to IV phenytoin load		
Valproate Depacon	IV: Up to 30 mg/kg IV (max rate 10 mg/kg/min) PO: (Depakote ER): 30 mg/kg ^{10,11, 12}	First Line: IV: 40 mg/kg (over 5-10 min); Max 3000 mg	Transient local irritation at injection site; GI discomfort orally	Not Studied	Oral Depakote loading not preferred due to intolerable GI side effects ¹³
Topiramate Topamax	400-800 mg ¹⁴	Refractory SE: 400-800 mg PO	Decreased sodium bicarbonate	Not Studied	Monitor for DDI
*for Status Epilepticus please refer to UCMC Status Epilepticus Guidelines					

Clinical Evidence:

1. Purcell TB, McPheeters RA, Feil M, et al. Rapid oral loading of carbamazepine in the emergency department. Ann Emerg Med. 2007;50:121-126

2. Fisher RS, Sachdeo RC, Pellock J, et al. Rapid initiation of gabapentin: a randomized, controlled trial. Neurology. 2001;56:743-748

3. Biton V, Rosenfeld WE, Whitesides J, et al. Intravenous lacosamide as replacement for oral lacosamide in patients with partial-onset seizures. Epilepsia. 2008;49:418-424

4. Lardizabal DV, Morris HH, Hovinga CA, et al. Tolerability and pharmacokinetics of oral loading with lamotrigine in epilepsy monitoring units. Epilepsia. 2003;44:536-539.

5. Koubeissi MZ, Amina S, Pita I, et al. Tolerability and efficacy of oral loading of levetiracetam. Neurology. 2008;70:2166-2170

6. Wheless JW, Clarke D, Hovinga CA, et al. Rapid infusion of a loading dose of intravenous levetiracetam with minimal dilution: a safety study. J Child Neurol. 2009;24:946-951

7. Swadron SP, Rudis MI, Azimian K, et al. A comparison of phenytoinloading techniques in the emergency department. Acad Emerg Med. 2004;11:244-252

8. Gallop K. Review article: phenytoin use and efficacy in the ED. Emerg Med Australas. 2010;22:108-118.

9. Rudis MI, Touchette DR, Swadron SP, et al. Cost-effectiveness of oral phenytoin, intravenous phenytoin, and intravenous fosphenytoin in the emergency department. Ann Emerg Med. 2004;43:386-397

10. Limdi NA, Knowlton RK, Cofield SS, et al. Safety of rapid intravenous loading of valproate. Epilepsia. 2007;48:478-483

11. Ghaleiha A. Haghighi M. Sharifmehr M. Jahangard L. Ahmadpanah M. Bajoghli H. Holsboer-Trachsler E. Brand S. Oral loading of sodium valproate compared to intravenous loading and oral maintenance in acutely manic bipolar patients. Neuropsychobiology. 2014; 70(1):29-35

12. Miller B. Perry W. Moutier C. Robinson S. Feifel D. Rapid oral loading of extended release divalproex in patients with acute mania. Gen Hosp Psychiatry. 2005; 24(3): 218-221

13. Tesoro EP. Brophy GM. Cohen H. Valproic Acid. In: Cohen H. eds. Casebook in Clinical Pharmacokinetics and Drug Dosing. McGraw-Hill; Accessed December 29, 2020.

https://accesspharmacy-mhmedical-com.proxy.libraries.uc.edu/content.aspx?bookid=1514§ionid=88805824

14. Asadi-Pooya AA. Jahromi MJ. Izadi S. Emami Y. Treatment of refractory generalized convulsive status epilepticus with enteral topiramate in resource limited setting. Seizure. 2015;24:114-117