

Table S1. Effect of Prototype Antiseizure Compounds on Seizure Burden in the TMEV Model: Seizures Observed at the Time of Drug Injection.

Compound	Mechanism of Action	Mouse 6 Hz (44 mA) ED₅₀^a	Dose mg/kg, IP	% VEH Cumulative Seizure Burden
Vehicle				100 ± 24.7
Carbamazepine	<i>Na⁺ Channel Blocker</i>	38.2	40	61.7 ± 18.7
Lacosamide	<i>Na⁺ Channel Blocker</i>	12.9	13	18.3 ± 4.6**
Lamotrigine ^b	<i>Na⁺ Channel Blocker</i>	43.4	20	49.6 ± 10.3**
Phenytoin	<i>Na⁺ Channel Blocker</i>	44.6	20	15.6 ± 4.5**
Ezogabine	<i>K⁺ Channel Opener</i>	32.5	20	41.6 ± 13.4
Clonazepam	<i>GABA_A Receptor Modulator</i>	0.17	0.2	60.4 ± 12.6
Phenobarbital	<i>GABA_A Receptor Modulator</i>	35.3	35	1.7 ± 1.3****
Tiagabine	<i>GABA Reuptake Inhibition</i>	1.0	1.3	48.0 ± 7.6
Ethosuximide	<i>T-Type Ca⁺⁺ Channel Blocker</i>	271	300	112 ± 20.2
Gabapentin	<i>α2δ Ca⁺⁺ Chanel Modulator</i>	>500	350	9.8 ± 3.3****
Levetiracetam	<i>SV2A Ligand</i>	>1000	1000	11.3 ± 5.4****
Topiramate	<i>Mixed</i>	>300	300	57.2 ± 8.8
Valproic Acid	<i>Mixed</i>	239	239	0.6 ± 0.6****

Vehicle (0.5% methylcellulose)

^aMetcalf et al., 2017

^bLamotrigine was administered once daily (QD), whereas other compounds noted were administered twice daily (BID).

*P<0.05, **P<0.01, ****P<0.0001 compared to VEH seizure burden from the same testing cohort; Mann-Whitney U test.

The vehicle-treated group was in the same testing cohort as topiramate (300 mg/kg) and ethosuximide (300 mg/kg) and was representative of other vehicle treatment groups.