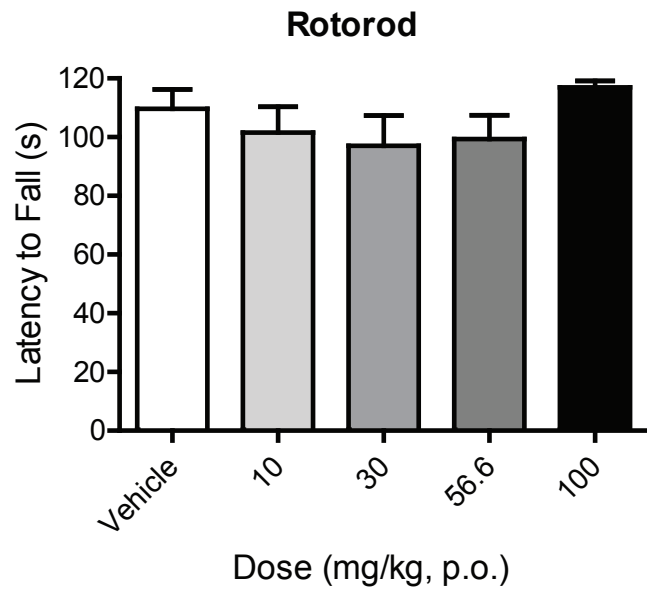
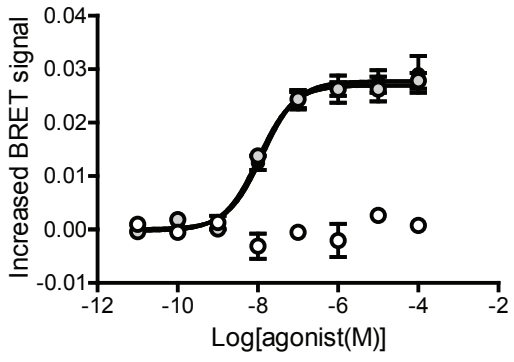


Supplemental Figure 1. VU0418506 does not induce deficits in rotarod performance at doses up to 100 mg/kg. Increasing doses of VU0418506 were administered orally to rats and their performance in a rotarod task was assessed as described in Materials and Methods. Data represent Mean \pm SEM from 12-15 animals.

Supplemental Figure 2. The CODA-RET signal is specific to the complemented pair. mGlu₂ was labeled with either the L1 fragment of luciferase or the L2 fragment. These constructs were transfected together and stimulated with the group II orthosteric agonist DCG-IV and a BRET signal was measured (black symbols). Co-transfection of unlabeled mGlu₄ did not change the DCG-IV response (gray symbols). Stimulation with the group III orthosteric agonist L-AP4 also did not induce a response (white symbols), indicating that activation is only read out if a receptor is complemented, and demonstrating that there is no contamination of the response by higher order complexes. Data represent three independent determinations performed in triplicate.



Supplemental Figure 1



Receptors	Agonists
● mGlu ₂ _L1+mGlu ₂ _L2	DCG IV
○ mGlu ₂ _L1+mGlu ₂ _L2+mGlu ₄	DCG IV
○ mGlu ₂ _L1+mGlu ₂ _L2+mGlu ₄	L-AP4

Supp Figure 2