

I. Key Concepts

- Major excitatory NT in brain: **Glutamate**
- Major inhibitory NT in brain: **GABA**
- Glutamate receptors have 2 large families:
 - Ligand-gated ion channels aka "ionotropic receptors"
 - G protein-coupled receptors called "metabotropic receptors"
- Ionotropic glutamate receptors divided into 3 classes which are named after the synthetic ligands that activate them:
 - **AMPA**
 - **Kainate**
 - **NMDA**
- Individual excitatory synapses typically express several different subtypes of ionotropic glutamate receptors as well as metabotropic receptors
- **NMDA receptors have 2 important biophysical properties (in contrast to AMPA and kainate):**
 - **Highly permeable to Calcium**
 - **Voltage dependent**
 - **ALLOWs CALCIUM ENTRY ONLY IF THE CELL IS DEPOLARIZED**
- AMPA receptors mediate vast majority of excitatory synaptic transmission in brain
 - NMDA receptors play important role in triggering synaptic plasticity and, when over-activated, in triggering excitotoxicity
- 8 subtypes of metabotropic glutamate receptors
 - **When localized to presynaptic terminal, they inhibit NT release**
 - When localized to the postsynaptic membrane, they exert complex modulatory effects thru specific signal transduction cascades, which can lead to **excitatory or inhibitory effects**
- Most studied form of synaptic plasticity: Long-term potentiation (LTP) in hippocampus
 - Triggered by strong activation of NMDA receptors and the consequent **large rise in postsynaptic Ca^{2+}**
 - There are several other types of LTP in the NS and are mediated via different mechanisms
- LTD = long-term depression
 - Long-lasting decrease in synaptic strength
 - Also occurs at most excitatory and some inhibitory synapses in brain
- 2 major classes of GABA receptor

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