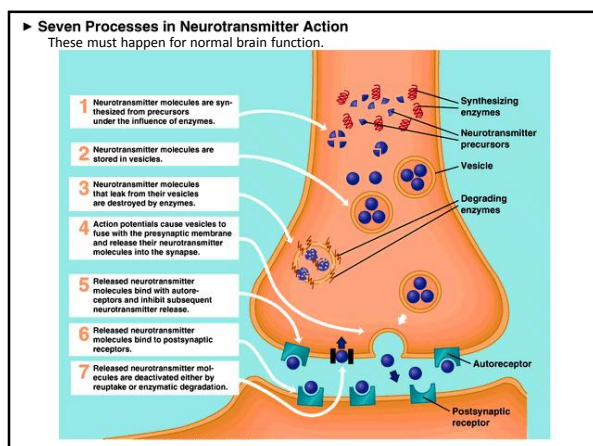


Cleaning Up Neurotransmitter

- Transmitter must be removed from synapse after its release & activation of receptors.
- Most neurotransmitters are recycled thru a process called "**reuptake**" – transported back into axon ending to be used again
- Less often an enzyme breaks down transmitter into inactive components.
- Clean up is critical to normal nervous system function.



• The "Big Seven"

(Best Known Neurotransmitters That our Psychoactive Drugs Will Influence)

- Acetylcholine (ACh)
 - Norepinephrine (NE)
 - Dopamine (DA)
 - Serotonin or 5-Hydroxytryptamine (5HT)
 - GABA
 - Endorphin
 - Glutamate
- <http://www.uni.edu/walsh/neurotransmitters.html>

•Acetylcholine (ACh)

- neurons using ACh =“cholinergic neurons”
- Where do you find them?
 - Nerves to skeletal muscle
 - Parasympathetic N.S.
 - Learning and memory areas of brain
- some drugs (Cognex (tacrine), Aricept, Reminyl) increase Ach actions
- others (“anticholinergics”) block its action

•Norepinephrine (NE)

- Where do you find NE neurons?
 - **Sympathetic N.S.**
 - **Brain areas involved in appetite, arousal, mood**
- Some drugs activate NE receptors (“sympathomimetics”)
- others block NE receptors (e.g. “beta-blockers”)

•Dopamine (DA)

- Very closely related to NE (“catecholamines”) and both NE & DA loosely related to serotonin (“monoamines”)
- Where is it found?
 - **basal ganglia (motor control)**
 - **limbic system (emotion, mood and “reward”)**
 - **frontal cortex (judgment & reasoning)**
 - **hypothalamus link to pituitary gland (hormone control)**
- some drugs increase DA (l-dopa)
- others block DA (antipsychotics)

•Serotonin or 5 Hydroxytryptamine (5HT)

- found in
 - **sleep & pain suppression areas of brain,**
 - **in limbic system (mood)**
 - **in sensory processing areas**
- several drugs increase 5HT (antidepressants)
- 5HT blockers are used to decrease nausea (e.g. Zofran)

•GABA

- best known inhibitory transmitter
- widely distributed in CNS
- seems necessary to keep neuron activity “in check” - without enough of it you might suffer from excessive neural activity causing anxiety or epilepsy
- several drugs increase the effects of GABA (benzodiazepines, alcohol)

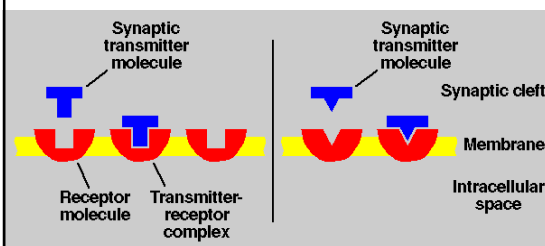
Glutamate

- Amino acid which acts as an excitatory transmitter almost everywhere in the CNS
- PCP blocks some glutamate receptors

•Endorphins

- Peptide family of transmitters which decrease pain perception and elevate mood
- Narcotic analgesic drugs act on endorphin receptors

Post-Synaptic Receptor Specificity

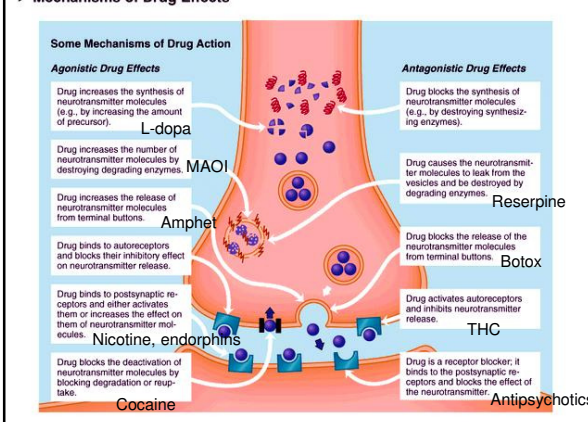


- Each transmitter binds only to its receptors, but there are multiple types of receptors for each.
- Drugs may increase or decrease the synaptic action of neurotransmitters. Drugs may affect only certain types of a transmitter's receptors.

•Ways Drugs May Affect Neurons

- Drugs may affect any of the normal neuron processes
 - Production of transmitter
 - Storage of transmitter in vesicles
 - Release of transmitter
 - Binding & action of transmitter at receptor sites
 - Elimination of transmitter by reuptake or enzymatic breakdown
- Drugs exert these actions by binding to proteins involved in these functions.
- The study of how drugs exert their effects on cells is "pharmacodynamics"

► Mechanisms of Drug Effects



•Drugs & Receptor Sites

- **AGONIST:** A drug that triggers or increases the usual synaptic effects of a transmitter
 - E.g. A drug which fits post-synaptic receptor sites and MIMICS action of transmitter
- **ANTAGONIST:** A drug prevents or decreases the usual synaptic effects of a transmitter
 - E.g. A drug which fits receptor site but **does not** trigger a response. This drug is a **BLOCKER**.

•Drug Actions

- **AGONIST** examples:
 - Narcotic pain relievers fit and activate opiate receptors **mimicking** the action of normal endorphin.
 - Nicotine fits into & stimulates ACh receptor sites, arousing the cortex like ACh
- **ANTAGONIST** examples:
 - Haldol **blocks** DA receptors in schizos
 - Naloxone **blocks** opiate receptors
 - Atropine & curare (discussed earlier) block different types of ACh receptors