

Monoamine transmitter formed from tryptophan (5hydroxytryptophan) Plays a major role in sexuality, depression (with elevated risk of suicide), bipolar disorder, and anxiety

- Disruption affects the suprachiasmatic nucleus (circadian rhythm)
- Predominantly produced in the raphe nuclei, of which the caudal raphe nuclei project to the medulla and spinal cord and play a role in the regulation of pain
- The rostral raphe nuclei project to the limbic system and cerebral cortex, and serotonin here is colocalized with norepinephrine





Raphe nuclei



Raphe nuclei

Brainstem 🚼



Raphe nuclei

anglia 🕂

Brainstem 🚼



Hypothalamus

ganglia

Raphe nuclei

Brainstem 🔡

Frontal cortex

ganglia •>

Limbic system

Hypothalamus

Raphe nuclei

Brainstem 🔡

Serotonin Receptors

- 5-HT1 Brain, intestinal nerves Neuronal inhibition, behavioral effects, cerebral vasoconstriction
- 5-HT2 Brain, heart, lungs, smooth muscle control, GI system, blood vessels, platelets - Neuronal excitation, vasoconstriction, behavioral effects, depression, anxiety
- 5-HT3 Limbic system, ANS Nausea, anxiety
- 5-HT4 CNS, smooth muscle Neuronal excitation, GI
- 5-HT5, 6, 7 Brain Depression

- First neurotransmitter described
- Derived from acetyl + choline
- Important role in myasthenia gravis at the neuromuscular junction
- Decreased in Alzheimer disease (may be partially reversed with rivastigmine)
- Pentameric receptor in muscles (two alpha, one beta, delta, and gamma subunits), tetrameric in CNS (two alpha, two beta)





Nucleus basalis of Meynert

Nucleus basalis of Meynert

Hippocampus and Amygdala

Frontal cortex

Nucleus basalis of Meynert

Hippocampus and Amygdala

Neocortex

Frontal cortex

Nucleus basalis of Meynert

Hippocampus and Amygdala

Acetylcholine Receptors

- M1 Nerves CNS excitation, gastric acid secretion
- M2 Heart, nerves, smooth muscle Cardiac inhibition, neural inhibition
- M3 -Glands, smooth muscle, endothelium Smooth, muscle contraction, vasodilation
- M4 CNS Not known
- M5 CNS Not known
- NM Skeletal muscles neuromuscular junction -Neuromuscular transmission
- NN Postganglionic cell body dendrites Ganglionic transmission

Acetylcholine Receptors



Not a Simple Picture

Signaling Paradigm

- Neurotransmitter signaling is not simply a one neurotransmitter one response paradigm.
- Instead, it consists of multiple neurotransmitters each acting through multiple second messenger pathways to create immediate effects and longterm potentiation.
- The time course and frequency of stimulation affects the degree of long-term potentiation and synaptic modeling.



PHOSPHOINOSITIDE SIGNALING PATHWAY



Neural mechanisms of synaptic plasticity, short and long-term memory

