Intro to Cognitive Neuroscience

Organization of the brain Some neurotransmitters

Some anatomical terms

- Dorsal
- Ventral
- Anterior
- Posterior
- Medial
- Lateral

Brain organization labels derive from development

- Neural tube forms three distinct bumps
- Forebrain most anterior of these
- Hindbrain most posterior of these
- Midbrain in-between
- In fully-developed human brain, these distinctions are much harder to see



Forebrain

- Divided into cerebral hemispheres (endbrain) and "between-brain" structures
- Cerebral hemispheres (telencephalon)
 - Cerebral cortex and connecting white matter

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• Subcortical: basal ganglia (motor), limbic system (emotion, learning)



Forebrain

- Divided into cerebral hemispheres (endbrain) and "between-brain" structures
- Between-brain (diencephalon)
 - Thalamus processes and distributes sensory and motor information
 - Hypothalamus maintains homeostasis; controls endocrine system; involved in emotional response.



Midbrain

- Tectum nuclei involved in visual and auditory systems.
- Substantia nigra and ventral tegmental area have large concentrations of dopaminergic cells.



Hindbrain

- Divided into cerebellum, pons, and medulla
- Cerebellum motor coordination, maintenance of posture
- Pons alertness, attention, aggression, emotion
- Medulla vital functions (heart rate, breathing, digestion, blood pressure, etc)

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Neurotransmitters

- Criteria for a substance being a neurotransmitter:
 - 1. Exists in presynaptic axon terminals
 - 2. Presynaptic cell contains enzymes for synthesizing substance
 - 3. Substance is released in significant quantities when nerve impulses reach terminals
 - 4. Receptors specific to substance on post-synaptic membrane
 - 5. Application of the substance causes post-synaptic potentials
 - 6. Blocking release of substance prevents pre-synaptic impulses from affecting post-synaptic potentials

Acetylcholine (ACh)

- First NT to be discovered, by Otto Loewi in 1921.
- Experiment design came to him in a dream.
- Showed that nerve stimulation => release of chemical, which affects other cells.
- Loewi called it "Vagusstoff" (vagus-stuff).





Acetylcholine

- NT at neuromuscular junction
- In brain, basal forebrain cholinergic system (BFCS) innervates cortex, hippocampus, limbic system.
- Blocking ACh in BCFS interferes with learning tasks.



Acetylcholine - nicotinic receptors

- ACh has two types of receptors
- Nicotinic receptors are ionotropic.
 Channel for Na⁺ and Ca²⁺, so excitatory.
- Receptor at neuromuscular junction; also some in CNS.
- Also enhance release of NT when located on terminals.





Acetylcholine - muscarinic receptors

- Muscarinic receptors are metabotropic.
- (Metabotropic receptors cause long-lasting changes in the post-synaptic cell, usually by activating a second-messenger system.)



Muscarinic receptor (slow, metabotropic)

Acetylcholine - muscarinic receptors

- Muscarinic receptors are metabotropic.
- (Metabotropic receptors cause long-lasting changes in the post-synaptic cell, usually by activating a second-messenger system.)
- Cortex, hippocampus, thalamus, striatum, and basal forebrain all have lots of muscarinic receptors.
- Muscarinic receptors are involved in cognitive and motor functions of ACh.

Norepinephrine (NE) (British: Noradrenaline (NA))

- Noradrenergic cells primarily located in pons and medulla.
- Cells from locus coeruleus in pons project to cortex, limbic system, thalamus, hypothalamus.
- Locus coeruleus cells involved in vigilance alertness to stimuli.
- NE also acts as a hormone.



Norepinephrine

- Norepinephrine receptors are metabotropic.
- Found in cortex, thalamus, hypothalamus, cerebellum, hippocampus, and amygdala.
- Four types of NE receptors. All activate second-messenger systems to cause changes w/ in the post-synaptic neuron.



Dopamine (DA)

- Dopaminergic cells primarily located in midbrain; two pathways.
 - Nigrostriatal path: cells in substantia nigra project to striatum
 - Mesolimbocortical path: cells in ventral tegmental area project to cortex and to limbic structures.
- Nigrostriatal path involved in motor control
- Mesolimbocortical path involved in reward and addiction



Image courtesy of the National Institutes of Health

Dopamine

- DA receptors are metabotropic
- Five types of DA receptors, classified as D₁-like and D₂-like
- D₁-like cause an increase in cAMP, D₂-like cause a decrease in cAMP
- DA and NE are very chemically similar; both are reuptaken from the synapse by similar transporter proteins.



Serotonin (5-HT)

- Serotonergic cells concentrated in raphe nuclei in brainstem
- Project to cortex, hippocampus, basal ganglia, limbic system
- 5-HT implicated in sleep, mood, anxiety



Image courtesy of the National Institutes of Health.

Serotonin

- 5-HT is reuptaken by the 5-HT transporter
 - SSRI antidepressants (like Prozac) block this transporter
- >15 types of 5-HT receptors
- Most are metabotropic (but 5-HT3 receptors are ionotropic and excitatory)



Glutamate

- The transmitter for fast excitatory transmission
- Ionotropic receptor types include AMPA and NMDA
- Metabotropic receptors work by a variety of pathways inhibit cAMP formation, activate phosphinositide system, inhibit glutamate release.



Glutamate

- High levels of glutamate can be toxic to cells.
- So, uptake of extracellular glutamate is important!
- Astrocytes as well as neurons have proteins for glutamate uptake.



A blood vessel covered by astrocytes (in green). Image courtesy of Zerd.

GABA (γ-aminobutyric acid)

- Workhorse inhibitory transmitter in the brain.
- 10%-40% of nerve terminals in cortx, hippocampus and substantia nigra are GABAergic.
- In cortex and hippocampus, lots of local GABAergic interneurons.
- GABAergic neurons from striatum to substantia nigra are projection neurons





- GABA_A receptors are ionotropic; allow Cl⁻ to flow into cell.
- GABA_B receptors are metabotropic inhibit formation of cAMP, stimulate K⁺ channels opening.

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