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# Behavioral Neuroscience Notes

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## PSYC 360

*Start*

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**Behavioral Neuroscience Lecture  
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## 7.1 Overview – Unit 3

- Neurotransmitters and Neuromodulators
  - Psychopharmacology
  - Disorders
  - Pain

### 7.1.1 Psychopharmacology in Detail

- The scientific study of the effects of drugs on the nervous system and behavior.
- Psychopharmacology is the study of how drugs affect the mind and behavior.
  - Psychotherapeutic drugs
  - Better understanding of how things normally work.

#### Principles of Drug Action

- *Selective Action* – Drugs are selective in their action on the nervous system.
  - *Sites of Action* – The location at which a drug interacts with the body to produce its effects.
  - Side effects are often due to the drug acting on sites other than the intended target.
    - Thus, side effects are relative to what our preferred site of action is.
  - *Example*: Opioids primarily affect the opioid receptor system.
- Drugs don't CREATE effects, they *modulate* ongoing cellular activity.
  - That is, they affect behavior by affecting neural transmission in some way.
  - *Agonist* – A drug that mimics or enhances the **effects of a neurotransmitter**.
    - Facilitates post synaptic effects.
    - *Example*: Morphine mimics endorphins, which are natural painkillers.
  - *Antagonist* – A drug that blocks or inhibits the effects of a neurotransmitter.
    - Inhibits post synaptic effects.
    - *Example*: Naloxone blocks the effects of opioids, reversing their effects.
  - Agonistic effects can become antagonistic if the drug is taken in excess.
    - *Example*: I make a neuron fire a neurotransmitter, but I also block the reuptake of that neurotransmitter.



## Basic Process

- *Precursor* – A substance from which another substance is formed. AKA, the ingredients used to make a neurotransmitter.

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- *Synthesis* – The process of creating a neurotransmitter from its precursors.
- Sometimes, we break down the precursor to build the neurotransmitter.

### 7.1.2 Other Ways of Agonist and Antagonist

- Block  $\text{Ca}^{2+}$  channels from opening (antagonist)
- *Mimetic* – Mimics the action of a neurotransmitter.
  - *Direct Agonist* – Binds to the same receptor as the neurotransmitter and mimics its effects.
  - *Indirect Agonist* – Binds to a different site on the receptor and enhances the effects of the neurotransmitter.
- Blocking agent
  - Competitive
    - *Direct Antagonist* – Binds to the same receptor as the neurotransmitter and blocks its effects.
  - Non-competitive
    - *Indirect Antagonist* – Binds to a different site on the receptor and blocks the effects of the neurotransmitter.
    - *Inverse Agonist* – Binds to the same receptor as the neurotransmitter and produces the opposite effect.
- *Depolarizing* or *Desensitizing Agent* – A drug that causes the AP to stay in a depolarized state; refusing to let the neuron go through another AP, and it stays in the absolute refractory period. (Antagonist)
- Interfere with vesicles (leaky or transporter proteins). (Antagonist)
- Interfere with docking proteins. (Antagonist)
- Selectively deactivate autoreceptors. (Agonist)
- Selectively activate autoreceptors. (Antagonist)



## 7.2 What Do These Chemicals Between Neurons Do?

- Transmit information.
  - Glutamate
  - GABA
  - Glycine
- Modulate information.
  - Every other neurotransmitter.

## 7.3 Classes of Neurotransmitters (Revisited from Lab)

- Amino Acids
  - Glutamate, GABA, Glycine
    - *Glutamate* – Synthesized from precursor glutamine by an enzyme called *glutaminase*. It is the most common excitatory neurotransmitter in the brain.
      - Related closely with the *NMDA* receptor, which is a type of glutamate receptor that is important for synaptic plasticity and memory formation.
      - One drug that binds to this site, *Phencyclidine (PCP)* (direct antagonist), is a drug that blocks the NMDA receptor and causes hallucinations and dissociation. Another drug that is thought to bind here, *Ketamine* (direct antagonist), is a dissociative anesthetic that is used in surgery and is also being studied as a treatment for depression.
    - **Reuptake and Deactivation**
      - Reuptake is done by the *excitatory amino acid transporters (EAATs)*. These are important because it reduces the change of excitotoxicity, which is believed to be involved in damage to the brain in stroke and amyotrophic lateral sclerosis (ALS, Lou Gehrig's disease).
    - *GABA* – Synthesized from precursor glutamate by an enzyme called *glutamic acid decarboxylase (GAD)*. It is the most common inhibitory neurotransmitter in the brain.
  - *Amines* (monoamines) – Derived from amino acids
    - Catecholamines
      - Contain catechol and derived from the amino acid tyrosine.
      - *Tyrosine* – Precursor for the catecholamines.
      - Dopamine (DA), Norepinephrine (NE), Epinephrine (Adrenaline)
      - Dopaminergic, Adrenergic, and Noradrenergic systems.
    - *Indolamines*



- *Serotonin* (5-HT)
- *Melatonin*
- Peptides (AKA: Neuropeptides)
  - Endogenous Opioids
- Acetylcholine (ACh)
- Lipids
  - *Anadamide* (Sanskrit for “bliss”) – Endogenous cannabinoid.
  - These appear to be synthesized on demand; produced and released as needed and not stored in synaptic vesicles.
  - Anadamide is deactivated by the enzyme *fatty acid amide hydrolase (FAAH)*.
- **Two Other Classes**
  - Nucleosides
    - Adenosine
  - Soluble Gases
    - *Nitric Oxide (NO)* – Required for an erection.

## 7.4 Acetylcholine (ACh)

- First neurotransmitter discovered.
- Otto von Loewy – Discovered ACh in 1921.
  - This guy took a frog heart and put it in saline. Then, he took simulated the parasympathetic part of the vagus nerve, and saw that the heart slowed down.
  - He then took the saline and put it in a different frog heart, and saw that the heart slowed down again.
  - “Vagusstoff” (ACh) – The chemical that was released from the vagus nerve that slowed the heart down.
  - Cholinergic – Referring to ACh.
- **Some Functions**
  - **Function in the ANS:**
    - Sympathetic
      - Spinal nerve leaves the cord and synapses in the paravertebral ganglion (ACh)
      - Then makes neuromuscular junction with smooth muscles and glands (NE)
      - *Neuromuscular Junction* – The synapse between a motor neuron and a muscle fiber.



- *Paravertebral Ganglion* – A ganglion located next to the spinal cord.
  - Except sweat glands (ACh)
- *Sympathetic Chain* – A chain of ganglia that runs parallel to the spinal cord. This is the reason for when you get anxious, ALL of your body gets anxious.
- Parasympathetic
  - Spinal nerve leaves the cord and synapse in the parasympathetic ganglion (ACh)
  - Then makes neuromuscular junction with smooth muscles and glands (ACh)
- The only NT in the parasympathetic branch.
- NT of the preganglionic sympathetic branch.
- **Function in the Somatic NS**
  - Excites the neuromuscular junction (ACh)
  - So, ACh is important for getting motor messages out to all kinds of muscles and glands.
- **Function in the CNS**
  - ACh is important in:
    - Learning and alertness (*Basal Forebrain*)—activates the cortex and facilitates learning.
    - *Nucleus Basalis* – Projects to the cortex
    - *Medial Septal Nucleus* and *Nucleus of Diagonal Band* – Projects to the hippocampus through the fornix.
    - Memory (*medial septal nucleus*)—modulate the hippocampus
    - REM sleep generation (*Pedunculopontine nucleus (PPT)* and *Laterodorsal Tegmental Nucleus (LDT)*)—projects to the pons and thalamus.
    - Reward system.
- **Synthesis and Metabolism**
- **Drugs and Disorders**

### 7.4.1 ACh Synthesis and Metabolism

- **Synthesis**
  - In a nutshell: A breakdown of lipids leads to Choline, which is the precursor for ACh. Acetate is the anion in vinegar (Acetic acid). Then, this is combined with Acetate to make ACh.
  - In more detail:
    - CoA attaches to an acetate ion (*Acetylcoenzyme A (acetyl-CoA)*).





- Then, *choline acetyltransferase (ChAT)* transfers the acetate from the acetyl-CoA to the choline molecule.
- Mnemonic: **ChAT**: From right to left: Transfers acetate to choline.
- **Metabolism**
  - ACh is broken down by the enzyme *acetylcholinesterase (AChE)* into acetate and choline. Nice and simple!
  - The choline is taken back up by active transport and reused, and the acetate is broken down and eliminated.

### 7.4.2 Two Types of Cholinergic Receptors

- Nicotinic Receptors
  - Agonist at low doses, but antagonist at high doses.
  - Iontropic.
  - Found at the Neuromuscular Junction in the PNS.
  - *Curare* (direct antagonist) – A drug that blocks nicotinic receptors, causing paralysis.
    - Competitive blocking agent
    - Paralysis, surgery
- Muscarinic Receptors
  - Comes from a hallucinogenic mushroom (*Amanita muscaria*).
    - **Don't confuse with Serotonin's Mescaline: Cactus; nor Psilocybin: Mushroom.**
  - Vikings (probably took this drug before raiding) and Koryaks (Nordic people who used this mushroom in religious practices).
  - Metabotropic receptors.
  - Predominates in the CNS (although, both types are found in the CNS).
  - *Atropine* (direct antagonist) – A drug that blocks muscarinic receptors, causing pupil dilation and increased heart rate.
    - Competitive blocking agent
    - Belladonna alkaloids (deadly nightshade)

## 7.5 MORE Drugs and Toxins Affecting ACh

- *Botulinum Toxin* – A waste product of *Clostridium botulinum*, which are bacteria who grows without oxygen.



- Interferes with  $\text{Ca}^{2+}$  influx channels, preventing the release of ACh.
- Because Botox causes paralysis, it can interfere with emotional *expression* because it paralyzes muscles like the orbicularis oculi.
- Additionally, since we know that expression influences experience, when we paralyze these muscles, then the emotional *experience* is also negatively affected.
- **Does Botox Decrease Emotional Experiences?**
  - Population: Women who want wrinkles gone.
  - One IV: two levels: Botox or restylane (dermal filler).
  - Method: Everyone had wrinkle reduction. AND, Everyone watches some emotion evoking movies.
  - Results: Botox group had less emotional experience than the restylane group.
  - **Is this a good thing?**
    - Another study takes a sample of depressed people and gives them either Botox or a placebo.
    - Results: 15% of placebo had a decrease in depression, while 52% of the Botox group had a decrease in depression.
- Botox can also be used to treat migraines, cerebral palsy, and hyperhidrosis (excessive sweating).
- *Black Widow Spider Venom* – A neurotoxin that causes the release of ACh at the neuromuscular junction, causing continual release of ACh and paralysis.
- *Cobra and Krait Venom* – A neurotoxin that blocks the binding of ACh to nicotinic receptors, causing paralysis.
- *AchE Blockers* – Comes into contact with the enzyme that breaks down ACh, causing an increase in ACh in the synaptic cleft.
  - Irreversible
    - Insecticides (Parathion)
    - Nerve gas: DFP (Diisopropylfluorophosphate (don't need to know the whole name)) and Sarin.
      - Readily crosses the blood-brain barrier so PNS and CNS are affected.
      - Antidote?
        - *Atropine* – A drug that blocks muscarinic receptors, preventing the effects of excess ACh.
        - *Pralidoxime* – A drug that reactivates AChE, allowing it to break down ACh again.
  - Reversible
    - *Neostigmine* (**Prostigmin**) and *Physostigmine* (**Antilirum**) – Drugs that inhibit AChE, increasing the amount of ACh in the synaptic cleft.
      - Doesn't cross the blood-brain barrier, so it only affects the PNS.



- Used to treat *myasthenia gravis* (a disease that causes muscle weakness and fatigue).
- Autoimmune disease that attacks nicotinic receptors at the neuromuscular junction.
- *Donepezil* (**Aricept**) and *rivastigmine* (**Exelon**) – These drugs do the same thing as the above drugs, but they cross the blood-brain barrier and are used to treat Alzheimer's disease and Parkinson's disease (only the cognitive part).

## 7.6 New Drug for Schizophrenia

- We'll talk about dopamine drugs later in this unit.
- This new drug now:
  - *Xanomeline and trospium chloride* (**Cobenfy**) – A drug that blocks the muscarinic receptors in the CNS, but not in the PNS.
  - Dopamine but also Ach!

## 7.7 Catecholamines

- Dopamine (DA)
- Norepinephrine (NE)
- Epinephrine (Adrenaline)

### 7.7.1 Dopamine (DA)

- Synthesis and Metabolism
- Function
- Drugs and Disorders

#### Dopamine Synthesis

- Tyrosine was first discovered from cheese (tyrosine = cheese).
  - Tyrosine is the precursor for DA, NE, and Epi.
  - *Tyrosine Hydroxylase* – The rate-limiting enzyme in the synthesis of catecholamines.
    - Converts tyrosine to L-DOPA.
    - L-DOPA is the precursor for DA, NE, and Epi.
  - L-DOPA is converted to DA by the enzyme *DOPA decarboxylase*.

## Dopamine Metabolism

- DA is broken down by the enzyme *Monoamine Oxidase (MAO)* into *Dihydroxyphenylacetic acid (DOPAC)*.
- Then, *Catechol-O-methyltransferase (COMT)* converts DOPAC into *Homovanillic acid (HVA)*.
- Also, starting from DA, we can use COMT to convert it to 3-methoxytyramine (3-MT), then with MAO, we can convert it to HVA.

## DA Function

- **Movement/Motor systems**
  - *Nigrostriatal System* – Starts in the substantia nigra and ends in the striatum (caudate nucleus and putamen).
  - Here's the route: We start at the striatum, which then sends an inhibitory GABA signal to the substantia nigra, who sends a reciprocal inhibitory DOPA signal back to the striatum nerve that sends an inhibitory GABA signal to the globus pallidus. Then, the globus pallidus excites the thalamus, who then excites the primary motor cortex, who then excites movement.
    - Note that if the inhibitory signal to the substantia nigra is limited, then the signal that the striatum sends to the globus pallidus is much stronger, which leads to a weaker signal to the thalamus, and thus to movements.
- Parkinson's Disease symptoms:
  - Weakness,
  - Tremor at rest,
  - Muscle rigidity,
  - Problems with balance,
  - Abnormal gait,
  - Trouble learning
- Treatment
  - *Reserpine (Raudixin)* for ↓ BP (Not in use anymore because it caused Parkinson's-like symptoms)
    - 1960's
      - Blocks monoamine transporters
        - Developed Parkinson's symptoms
        - Can't fill vesicles and DA is lowered
      - Then, discovered Substantia Nigra was pale.
    - L-DOPA can be a direct treatment for Parkinson's as well.
    - *MPTP* – Neurotoxin for DA cells in the Nigrostriatal System (which is not endogenous).



- **History of MPTP – or why you shouldn’t use illicit drugs**
  - 1982 – young California heroin users
  - Had used what they THOUGHT was synthetic heroin
    - *MPPP* – Opioid analgesic drug
      - Not used clinically
      - Illegally manufactured for recreational drug use
    - INSTEAD it was MPTP (oh no!)
      - They instantly developed Parkinson’s-like symptoms
      - Bad for them, but good for us because we can study it.
        - Led to animal model development and possible treatment ideas.
  - We don’t know why Parkinson’s patient’s cells are dying, but maybe something similar.
  - MPTP is converted to the chemical *MPP+* by the enzyme MAO (which is what breaks down DA), which is what damaged the cells.
    - Question: Could MAO-I improve Parkinson’s?
    - Yes!
    - *Deprenyl*, also called *selegiline* (**Eldepryl, Jumex**) – A drug that inhibits MAO, can slow down progression of the disease.
- New treatment
  - Molecule keeps proteins from misfolding
  - *Lewy Bodies* – Misfolded proteins that are found in the brains of people with Parkinson’s.
    - These are toxic to DA cells
- *Huntington’s Chorea* – A genetic disorder that leads to uncontrolled movements and cognitive decline.
  - Too little GABA from the Striatum to the Substantia Nigra causes an increase in dopamine back to the Striatum which, in turn, lessens the signal to the Globus Pallidus, which increases overall movements.
  - *Tetrabenazine* (**Xenazine**) – Drug that inhibits the DA vesicle transporters.
  - *Pallidotomy* – A surgery that affects the Globus Pallidus to inhibit movement.
- *Choreoathetotic Movements* – too much movement
  - *Athetosis* – Slow continually writing movements
  - *Choreic* (to dance) – Rapid, purposeless, involuntary movements
- **Behavioral Arousal and Attention**
  - Narcolepsy
    - *Methylphenidate* (**Ritalin**) – A drug that increases DA and NE in the brain, used to treat ADHD, but can also be used for narcolepsy.



- *Hypocretine* – A neuropeptide that is involved in the regulation of sleep and wakefulness.
  - Created by the lateral hypothalamus.
  - Hypocretine: *Hypo* for *hypothalamus*, *cretine* for *secretin* (a hormone).
  - *Orexin* – Another name for hypocretine; makes you want to eat.
- From hypocretine, researchers developed an antagonist for the orexin receptor, which is used to treat insomnia. This drug is called *Suvorexant* (**Belsomra**).
- **Treatment**
  - *TAK-994* – OX2R (Orexin-2 receptor) Agonist
  - *Hcrt-1* – Intranasal hypocretine-1 (orexin-1) agonist
  - Hypocretine Cell Transplant
  - Gene Therapy: *introduce* preprohypocretin gene into the brain to make more hypocretine.
  - Opiates (exogenous) can increase the number of hypocretin-producing cells in the brain.
  - Indirect role for opiate agonists in treating narcolepsy.
- ADHD
  - Uses Methylphenidate for selective attention.
- *Mesocortical System*
  - From ventral tegmental nucleus to prefrontal cortex, limbic CORTEX, hippocampus, all frontal lobes, and association areas of parietal and temporal lobes in primates.
  - Short-term memories, planning, and problem-solving are all associated with this system.
- **Reinforcement and Reward**
  - *Mesolimbic System (MLS)* – Responsible for reward and reinforcement.
    - From ventral tegmental nucleus to limbic system
      - Amygdala, hippocampus, and nucleus accumbens.
      - Opioids cause the release of dopamine at the nucleus accumbens, which is the pleasure center of the brain.
  - James Olds & Peter Milner (1954)
    - They asked: “Does electrical stimulation of the reticular formation facilitate learning?”
    - James Olds visits a conference and listens to Neal Miller, who says electrical stimulation is aversive, so it should be avoided.
    - One lone rat was put in a box with a lever, and when the rat pressed the lever, it would get a shock to the reticular formation. He ended up pressing the lever 700 times per hour.



- More studies of this
  - Skinner box
    - Rats press 2000 times per hour for a shock to the MLS.
    - Monkeys press 8000 times per hour for a shock to the MLS.
    - Starving animals will choose the MLS over food 80% of the time.
    - They also press the button for these conditions too:
      - Thirsty,
      - Getting shocked (at their feet),
      - Mother instincts.
- Delgado (1969) – For people who were getting their brain stimulated for seizures, this researcher also asked them about what they thought of the stimulation. They all thought that it was pleasurable.

## **Part II**

# **Behavioral Neuroscience Lab Notes**



## 8.1 What Do We Know About Sleep?

- Sleep is an active process.
- Sleepiness and alertness are controlled in part by a biological clock.
- Things can go wrong
  - There are whole books that discuss sleep disorders.
- *Suprachiasmatic nucleus (SCN)* is the master clock of the body.
  - The SCN is located in the hypothalamus and is responsible for regulating circadian rhythms.
  - Note the name of the SCN, it is *above* the *chiasm* of the optic nerve.
- Three kinds of rhythms:
  - *Ultradian* rhythms: cycles shorter than 24 hours (e.g., heart rate, respiration).
  - *Circadian* rhythms: cycles of about 24 hours (e.g., sleep-wake cycle, body temperature).
    - Our circadian rhythm is an endogenous clock that is influenced by exogenous factors.
    - *Free-running* is when the circadian rhythm is not influenced by external cues (e.g., light, temperature).
    - It is about 24.2 hours in humans.
  - *Infradian* rhythms: cycles longer than 24 hours (e.g., menstrual cycle, seasonal changes).
- *Zeitgeber* is a stimulus that helps to regulate the biological clock (e.g., light, temperature).
- Human clocks run long when left free running. Rats are short.



## 8.2 What is Sleep?

- For regular people, sleep is behaviorally defined as a state of reduced movement, species specific posture, reduced response to stimuli, and reversibility.
- For sleep researchers, they take a more physiological definition of sleep.
  - *Polysomnography (PSG)* is a method of recording various physiological signals during sleep, including:
    - **Electroencephalography (EEG)**: measures electrical activity in the brain.
    - **Electromyogram (EMG)**: measures muscle activity.
    - **Electrooculogram (EOG)**: measures eye movements.
  - Rechtschaffen and Kales (1968) defined sleep stages based on EEG patterns.
  - In 2007, the American Academy of Sleep Medicine (AASM) updated the sleep stage criteria.

Name	Frequency	Amplitude	Description	State
<b>Beta</b> $\beta$	12 – 50 Hz (variable)	Lower and Variable	Desynchronous	Awake and Paying Attention
<b>Alpha</b> $\alpha$	8 – 12 Hz	50 Microvolts	Synchronous	Relaxed Wakefulness (eye closed, not fully attending, and usually largest occipitally)
<b>Theta</b> $\theta$	3.5 – 7.5 Hz	Low in voltage Microvolts	Synchronous	Drowsy, Light Sleep (Stage 1)
<b>Delta</b> $\delta$	1 – 3.5 Hz	20 – 200 Microvolts	Synchronous	Deep Sleep (Stages 3 and 4)

Table 8.1: Summary of EEG Wave Characteristics

## 3.1

**Q3.1.1 Short Answer:** What is psychopharmacology, and why do we study it?

*Answer:*

**Q3.1.2 Multiple Choice:** Which of the following is NOT a function of psychopharmacology?

- (A) Study the effects of drugs on the nervous system
- (B) Study the effects of drugs on behavior
- (C) Study the effects of drugs on the immune system
- (D) Study the effects of drugs on neurotransmitter systems

**Q3.1.3 Fill in the Blank:** The location at which a drug interacts with the body to produce its effects is called the \_\_\_\_\_.

**Q3.1.4 Short Answer:** What is the difference between an agonist and an antagonist?

*Answer:*

**Q3.1.5 True or False:** Drugs directly create effects in the body.

*Answer:*

**Q3.1.6 Multiple Choice:** Which of the following is an example of a drug that acts as an agonist? (Some of these drugs we haven't talked about, so pick the one you *know* is an agonist.)

- (A) Naloxone
- (B) Morphine
- (C) Curare
- (D) Atropine

**Q3.1.7 Short Answer:** What is selective action?

*Answer:*

**Q3.1.8 Short Answer:** What is an example of how an agonistic effect can become antagonistic?

*Answer:*



**Q3.1.9 Multiple Choice:** What is a precursor?

- (A) A substance that inhibits neurotransmitter release
- (B) A substance that enhances neurotransmitter release
- (C) A substance from which another substance is formed
- (D) A substance that blocks neurotransmitter receptors

*Answer:*

**Q3.1.10 Fill in the Blank:** The process of creating a neurotransmitter from its precursors is called \_\_\_\_\_.

**Q3.1.11 Fill in the Blanks:** A(n) \_\_\_\_\_ agonist binds to the same receptor as the neurotransmitter and \_\_\_\_\_ its effects, while a(n) \_\_\_\_\_ agonist binds to a different site on the receptor and \_\_\_\_\_ the effects of the neurotransmitter.

**Q3.1.12 Fill in the Blanks:** A(n) \_\_\_\_\_ antagonist binds to a different site on the receptor and \_\_\_\_\_ the effects of the neurotransmitter, while a(n) \_\_\_\_\_ antagonist binds to the same receptor as the neurotransmitter and \_\_\_\_\_ its effects.

**Q3.1.13 Multiple Choice:** Drugs that cause the action potential to stay in a depolarized state are called:

- |                 |                         |
|-----------------|-------------------------|
| (A) Agonists    | (B) Depolarizing agents |
| (C) Antagonists | (D) Inverse agonists    |



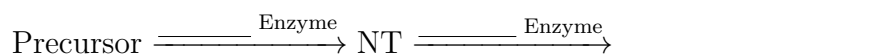
**Q3.1.14 Matching:** Match the following examples with them either being an antagonist or an agonist. (Some of these may be direct or indirect. Specify each one.)

### Choices

- (a) *Curare*
- (b) *Atropine*
- (c) *Morphine*
- (d) *Naloxone*
- (e) *Botulinum Toxin*
- (f) Interfering with docking proteins
- (g) Blocking the reuptake of a neurotransmitter
- (h) *Sarin*
- (i) Interfering with vesicles
- (j) Blocking receptors
- (k) Black widow spider venom
- (l) Cobra and Krait Venom
- (m) Parathion
- (n) DFP
- (o) *Physostigmine*

- (1) Direct antagonist ..... \_\_\_\_\_
- (2) Indirect antagonist ..... \_\_\_\_\_
- (3) Direct agonist ..... \_\_\_\_\_
- (4) Indirect agonist ..... \_\_\_\_\_
- (5) Antagonist ..... \_\_\_\_\_
- (6) Agonist ..... \_\_\_\_\_

**Q3.1.15 Fill in the Blanks:** In the following diagram, label the specific enzyme for each arrow, and identify the outcome:





**Q3.1.16 Long(-ish) Answer:** Describe the difference between a neurotransmitter and a neuromodulator.

*Answer:*



## 3.2

**Q3.2.1 Multiple Choice:** Which of the following neurochemicals does NOT transmit information (according to our notes)?

(A) Dopamine

(B) Glutamate

(C) GABA

(D) Glycine

**Q3.2.2 Fill in the Blank:** Peptides are short chains of \_\_\_\_\_.

**Q3.2.3 Fill in the Blanks:** The difference between opioids and opiates are that opioids are \_\_\_\_\_ and opiates are \_\_\_\_\_.

**Q3.2.4 Short Answer:** What is the pain pathway for the face? What about from the neck down? (Generally speaking.)

*Answer:*

**Q3.2.5 Fill in the Blanks:** The three types of opioid receptors are \_\_\_\_\_, \_\_\_\_\_, and \_\_\_\_\_.

**Q3.2.6 Long Answer:** What are each of the three opioid receptors responsible for, and what neurochemicals bind to each the most?

*Answer:*

**Q3.2.7 Multiple Choice:** Prostaglandins become active during

(A) Resting-and-Digesting

(B) Crying

(C) Daydreaming

(D) Bleeding

**Q3.2.8 Short Answer:** What are the functions of opioids? (List the main effects and the side effects.)

*Answer:*

**Q3.2.9 Fill in the Blank:** *Cyclooxygenase (COX)* is an enzyme that converts inactive \_\_\_\_\_ to its active state.



**Q3.2.10 Short Answer:** List the characteristics for the direct pain pathway and the indirect pain pathway.

*Answer:*

**Q3.2.11 Fill in the Blanks:** Pain arrives at the \_\_\_\_\_, then travels to the \_\_\_\_\_. Once there, it is processed by several brain regions. First, the \_\_\_\_\_ contributes to arousal. Then, the \_\_\_\_\_, particularly the anterior cingulate cortex (ACC), processes the emotional aspects of pain. When the pain is overwhelming, the \_\_\_\_\_ activates and releases endogenous opioids to reduce the sensation—this allows a person, for example, to escape danger despite a severe injury. Finally, the \_\_\_\_\_ and other areas help interpret and associate the pain with context.

**Q3.2.12 Matching:** Match the following drugs with their respective NSAID class.

**Choices**

- (a) Ibuprofen
- (b) Aspirin
- (c) Diflunisal
- (d) Naproxen
- (e) Salsalate
- (f) Ketoprofen

- (1) Proprionic Acid Derivatives ..... \_\_\_\_\_
- (2) Salicylates ..... \_\_\_\_\_

**Q3.2.13 True or False:** *Celecoxib* (**Celebrex**), a COX-2 Inhibitor, was removed from the market because it causes heart attacks and stroke.

*Answer:*

**Q3.2.14 Long Answer:** Some studies show that both the placebo effect and acupuncture can be blocked by Naloxone, an opioid antagonist. What does this suggest about the mechanism of acupuncture's pain-relieving effects? Does this prove that acupuncture is not entirely a placebo?

*Answer:*



**Q3.2.15 True or False:** The term *colocalized* means two or more neurotransmitters are released from two separate neurons at the same time.

*Answer:*

**Q3.2.16 Short Answer:** What is the definition of pain? (DO NOT say this exam!!!!!!!)

*Answer:*



### 3.3

**Q3.3.1 True or False:** The amines (monoamines) are derived from amino acids.

*Answer:*

**Q3.3.2 Short Answer:** Name three neurotransmitters that fall under the amino acid category.

*Answer:*

**Q3.3.3 Fill in the Blank:** The two indolamines are \_\_\_\_\_ and \_\_\_\_\_.

**Q3.3.4 Multiple Choice:** What amino acid are indolamines derived from?

(A) Tryptophan      (B) Tyrosine      (C) Glutamate      (D) Glycine

**Q3.3.5 Fill in the Blank:** The precursor to glutamate is \_\_\_\_\_, and the enzyme that synthesizes glutamate from it is \_\_\_\_\_.

**Q3.3.6 Short Answer:** What receptor does ketamine bind to, and what is its effect?

*Answer:*

**Q3.3.7 Fill in the Blank:** The enzyme \_\_\_\_\_ deactivates anandamide.

**Q3.3.8 True or False:** The most common excitatory neurotransmitter in the brain is GABA.

*Answer:*

**Q3.3.9 Fill in the Blank:** The drug \_\_\_\_\_ is a direct antagonist of the NMDA receptor and can cause hallucinations and dissociation.

**Q3.3.10 Short Answer:** What transporters are responsible for glutamate reuptake, and why is this process important?

*Answer:*

**Q3.3.11 Multiple Choice:** Which receptor is closely associated with glutamate and is important for synaptic plasticity and memory formation?

(A) GABA receptor      (B) NMDA receptor  
(C) Serotonin receptor      (D) Dopamine receptor





**Q3.3.12 Short Answer:** What enzyme converts glutamate into GABA, and what type of neurotransmitter is GABA?

*Answer:*

**Q3.3.13 Fill in the Blanks:** The three catecholamine neurotransmitters are \_\_\_\_\_, \_\_\_\_\_, and \_\_\_\_\_.

**Q3.3.14 Multiple Choice:** What do all catecholamines contain, and what amino acid are they derived from?

- (A) Catechol and are derived from tryptophan
- (B) Catechol and are derived from tyrosine
- (C) Indole and are derived from tryptophan
- (D) Indole and are derived from tyrosine

**Q3.3.15 Fill in the Blank:** The enzyme \_\_\_\_\_ converts tyrosine into L-DOPA.

**Q3.3.16 Short Answer:** Explain how botox interferes with emotional expression.

*Answer:*

**Q3.3.17 Fill in the Blank:** The orbicularis oculi muscle influences \_\_\_\_\_.

**Q3.3.18 True or False:** Tyrosine is the precursor for serotonin.

*Answer:*

**Q3.3.19 Short Answer:** What are the names of the systems that use dopamine, norepinephrine, and epinephrine? *Answer:*

**Q3.3.20 Fill in the Blanks:** Melatonin is synthesized from \_\_\_\_\_ and is involved in regulating \_\_\_\_\_.

**Q3.3.21 Short Answer:** What is another name for peptides in the context of neurotransmitters, and give an example. *Answer:*

**Q3.3.22 Multiple Choice:** What is the name of the endogenous cannabinoid neurotransmitter whose name means “bliss” in Sanskrit?

- (A) Anandamide
- (B) Cannabidiol
- (C) Tetrahydrocannabinol (THC)
- (D) 2-Arachidonoylglycerol (2-AG)

**Q3.3.23 Short Answer:** How are lipid-based neurotransmitters synthesized and stored?

*Answer:*

**Q3.3.24 Fill in the Blank:** The gaseous neurotransmitter that is required for an erection is \_\_\_\_\_.

**Q3.3.25 Short Answer:** Name one neurotransmitter that is a nucleoside. What is its function?

*Answer:*



**Q3.3.26 Fill in the Blanks:** Fill in the following spaces that describe the process of dopamine metabolism: DA is broken down by \_\_\_\_\_ into \_\_\_\_\_. Then, \_\_\_\_\_ converts it into \_\_\_\_\_.

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### 3.4:

**Q3.4.1 Short Answer:** What does cholinergic mean?

*Answer:*

**Q3.4.2 Multiple Choice:** Who first discovered acetylcholine in 1921?

- |                    |                  |
|--------------------|------------------|
| (A) Otto von Loewy | (B) James Olds   |
| (C) Neal Miller    | (D) Peter Milner |

**Q3.4.3 Short Answer:** What experiment led to the discovery of acetylcholine?

*Answer:*

**Q3.4.4 Fill in the Blank:** The original name given to acetylcholine by its discoverer was \_\_\_\_\_.

**Q3.4.5 Short Answer:** What are the two types of ACh receptors?

*Answer:*

**Q3.4.6 True or False:** Acetylcholine is the only neurotransmitter used in the parasympathetic branch of the autonomic nervous system.

*Answer:*

**Q3.4.7 Fill in the Blanks:** In the sympathetic nervous system, ACh is used at the \_\_\_\_\_, while NE is used at the \_\_\_\_\_.

**Q3.4.8 Multiple Choice:** Which of the following is NOT a function of ACh in the CNS?

- |                            |                     |
|----------------------------|---------------------|
| (A) Learning and alertness | (B) Memory          |
| (C) REM sleep generation   | (D) Pain modulation |

**Q3.4.9 Short Answer:** Describe the synthesis of acetylcholine.

*Answer:*

**Q3.4.10 Fill in the Blanks:** The precursor to acetylcholine is \_\_\_\_\_ and the enzyme that synthesizes acetylcholine is \_\_\_\_\_.

**Q3.4.11 Short Answer:** Explain how acetylcholine is metabolized.

*Answer:*



**Q3.4.12 Multiple Choice:** Which type of ACh receptor is ionotropic?

- (A) Nicotinic receptors
- (B) Muscarinic receptors
- (C) Both nicotinic and muscarinic receptors
- (D) Neither nicotinic nor muscarinic receptors

**Q3.4.13 Short Answer:** Explain what the sympathetic chain is, and where it is located.

*Answer:*

**Q3.4.14 Fill in the Blank:** The drug \_\_\_\_\_ is a direct antagonist of nicotinic receptors, causing paralysis.

**Q3.4.15 True or False:** Atropine blocks muscarinic receptors and is derived from the plant known as belladonna alkaloids (deadly nightshade).

*Answer:*

**Q3.4.16 Short Answer:** How does Botulinum Toxin interfere with acetylcholine function?

*Answer:*

**Q3.4.17 Fill in the Blanks:** Black Widow Spider venom causes \_\_\_\_\_ of ACh, while Cobra venom \_\_\_\_\_ ACh receptors.

**Q3.4.18 Multiple Choice:** Which of the following is a reversible AChE blocker used to treat myasthenia gravis?

- (A) Sarin
- (B) Parathion
- (C) Neostigmine (Prostigmin)
- (D) DFP (Diisopropylfluorophosphate)

**Q3.4.19 True or False:** *Donepezil* (**Aricept**) crosses the blood-brain barrier and is used to treat the cognitive symptoms of Alzheimer's disease.

*Answer:*

**Q3.4.20 True or False:** Nicotinic receptors are antagonists at low doses, but agonists at high doses.

*Answer:*

**Q3.4.21 Multiple Choice:** In the PNS, where are nicotinic receptors predominantly located?

- (A) Brain and spinal cord
- (B) Neuromuscular junctions
- (C) Autonomic ganglia
- (D) All of the above

**Q3.4.22 Long Answer:** Define the neuromuscular junction and the paravertebral ganglion.

*Answer:*



**Q3.4.23 Multiple Choice:** In the sympathetic nervous system, which neurotransmitter is used at the neuromuscular junction with smooth muscles and glands?

- (A) Acetylcholine (B) Norepinephrine  
(C) Dopamine (D) Serotonin

**Q3.4.24 True or False:** In the sympathetic nervous system, acetylcholine is the neurotransmitter used at the neuromuscular junction with sweat glands.

*Answer:*

**Q3.4.25 Fill in the Blanks:** The \_\_\_\_\_ is a chain of ganglia that runs parallel to the spinal cord. This is why when you get anxious, \_\_\_\_\_ of your body responds at once.

**Q3.4.26 Short Answer:** Compare the neurotransmitters used in the parasympathetic nervous system versus the sympathetic nervous system.

*Answer:*

**Q3.4.27 Multiple Choice:** Which of the following statements about acetylcholine in the autonomic nervous system is FALSE?

- (A) ACh is the only neurotransmitter in the parasympathetic branch  
(B) ACh is used at preganglionic synapses in both sympathetic and parasympathetic branches  
(C) ACh is used at postganglionic synapses to sweat glands in the sympathetic branch  
(D) ACh is the primary neurotransmitter at the neuromuscular junction with smooth muscles in the sympathetic branch

**Q3.4.28 Fill in the Blank:** In the somatic nervous system, ACh \_\_\_\_\_ the neuromuscular junction.

**Q3.4.29 Short Answer:** Explain the role of acetylcholine in the somatic nervous system.

*Answer:*

**Q3.4.30 Matching:** Match each brain structure with its projection target.

### Choices

- (a) Nucleus Basalis  
(b) Medial Septal Nucleus and Nucleus of Diagonal Band  
(c) Pedunculopontine nucleus (PPT) and Laterodorsal Tegmental Nucleus (LDT)

(1) Projects to the cortex ..... \_\_\_\_\_



- (2) Projects to the hippocampus through the fornix ..... \_\_\_\_\_
- (3) Projects to the pons and thalamus ..... \_\_\_\_\_

**Q3.4.31 Multiple Choice:** Which structure in the basal forebrain that uses ACh is primarily responsible for activating the cortex and facilitating learning?

- (A) Nucleus Basalis (B) Medial Septal Nucleus  
(C) Nucleus of Diagonal Band (D) Pedunculopontine nucleus

**Q3.4.32 True or False:** The Medial Septal Nucleus, which uses ACh, primarily modulates the amygdala.

*Answer:*

**Q3.4.33 Long Answer:** Explain the function of acetylcholine in REM sleep generation, including the specific brain structures involved.

*Answer:*

**Q3.4.34 Fill in the Blanks:** The \_\_\_\_\_ and \_\_\_\_\_ are structures that use acetylcholine and project to the hippocampus through the fornix.

**Q3.4.35 Short Answer:** What are the four main functions of acetylcholine in the central nervous system?

*Answer:*

**Q3.4.36 True or False:** When comparing the sympathetic and parasympathetic nervous systems, both use ACh at their preganglionic synapses.

*Answer:*

**Q3.4.37 Multiple Choice:** Which of the following correctly describes the neurotransmitter pathway in the parasympathetic nervous system?

- (A) ACh at preganglionic synapse, ACh at postganglionic synapse  
(B) ACh at preganglionic synapse, NE at postganglionic synapse  
(C) NE at preganglionic synapse, ACh at postganglionic synapse  
(D) NE at preganglionic synapse, NE at postganglionic synapse

**Q3.4.38 Shoet Answer:** What is the connection between vikings, Koryaks and ACh?

*Answer:*

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### 3.5:

- Q3.5.1 Fill in the Blanks:** The three catecholamines are \_\_\_\_\_, \_\_\_\_\_, and \_\_\_\_\_.
- Q3.5.2 Multiple Choice:** What is the precursor for dopamine?  
(A) Tyrosine                      (B) L-DOPA                      (C) Tryptophan                      (D) Choline
- Q3.5.3 Fill in the Blank:** The rate-limiting enzyme in the synthesis of catecholamines is \_\_\_\_\_.
- Q3.5.4 Short Answer:** Describe the pathway of dopamine synthesis from its amino acid precursor.  
*Answer:*
- Q3.5.5 True or False:** The word “tyrosine” is derived from a word meaning “tire.”  
*Answer:*
- Q3.5.6 Multiple Choice:** Which pathway is involved in movement and motor control?  
(A) Nigrostriatal system                      (B) Mesocortical system  
(C) Mesolimbic system                      (D) Tuberoinfundibular system
- Q3.5.7 Fill in the Blanks:** The nigrostriatal pathway starts in the \_\_\_\_\_ and ends in the \_\_\_\_\_.
- Q3.5.8 Short Answer:** List four symptoms of Parkinson’s disease.  
*Answer:*
- Q3.5.9 Multiple Choice:** What neurotoxin led to the development of an animal model for Parkinson’s disease?  
(A) MPTP                      (B) MPPP                      (C) MPP+                      (D) MAO
- Q3.5.10 Fill in the Blank:** The misfolded proteins found in the brains of people with Parkinson’s disease are called \_\_\_\_\_.
- Q3.5.11 True or False:** In Huntington’s Chorea, there is too much GABA from the Striatum to the Substantia Nigra.  
*Answer:*
- Q3.5.12 Long Answer:** Explain how the MPTP incident in 1982 contributed to our understanding of Parkinson’s disease.  
*Answer:*



**Q3.5.13 Fill in the Blank:** *Methylphenidate* (**Ritalin**) increases levels of \_\_\_\_\_ and \_\_\_\_\_ in the brain.

**Q3.5.14 Multiple Choice:** Which system is primarily responsible for reward and reinforcement?

- (A) Nigrostriatal system (B) Mesocortical system  
(C) Mesolimbic system (D) Tuberoinfundibular system

**Q3.5.15 Short Answer:** What neuropeptide, also called orexin, is involved in the regulation of sleep and wakefulness?

*Answer:*

**Q3.5.16 Fill in the Blank:** The drug \_\_\_\_\_ is an orexin receptor antagonist used to treat insomnia.

**Q3.5.17 True or False:** The mesocortical system is involved in short-term memory, planning, and problem-solving.

*Answer:*

**Q3.5.18 Multiple Choice:** Which researchers discovered that electrical stimulation of certain brain areas could be rewarding rather than aversive?

- (A) Otto von Loewy and Vagusstoff (B) James Olds and Peter Milner  
(C) Neal Miller and Delgado (D) Lateral hypothalamus researchers

**Q3.5.19 Short Answer:** What structure within the limbic system is considered the “pleasure center” of the brain?

*Answer:*

**Q3.5.20 Fill in the Blanks:** The following is a paragraph that describes dopamine synthesis: Tyrosine is converted to \_\_\_\_\_ by the enzyme \_\_\_\_\_. This converted form is then used to create dopamine by the enzyme \_\_\_\_\_.

**Q3.5.21 Multiple Choice:** Which of the following is NOT a function of dopamine in the CNS?

- (A) Movement and motor control (B) Reward and reinforcement  
(C) Learning and memory (D) Sleep-wake cycles

**Q3.5.22 Short Answer:** Describe the metabolism of dopamine.

*Answer:*

**Q3.5.23 Short Answer:** Define choreoathetotic movements.

*Answer:*



**Q3.5.24 Fill in the Blanks:** The term \_\_\_\_\_ refers to slow, continuous writhing movements, while \_\_\_\_\_ (from the Greek word for "dance") refers to rapid, purposeless, involuntary movements.

**Q3.5.25 True or False:** Both athetosis and choreic movements are characterized by too little movement.

*Answer:*

**Q3.5.26 Short Answer:** Where in the brain is hypocretin produced?

*Answer:*

**Q3.5.27 Multiple Choice:** Which drug increases both dopamine and norepinephrine in the brain and can be used to treat narcolepsy?

(A) Suvorexant (Belsomra)

(B) Methylphenidate (Ritalin)

(C) TAK-994

(D) Hypocretin

**Q3.5.28 Short Answer:** Explain the difference between athetosis and choreic movements.

*Answer:*

**Q3.5.29 Fill in the Blank:** \_\_\_\_\_ is a neuropeptide involved in the regulation of sleep and wakefulness that is also known as orexin.