

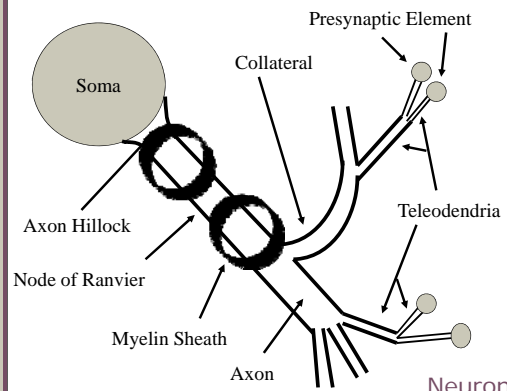


Overview of Neurons

Psychology 472

Pharmacology of Psychoactive
Drugs

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Structures

Soma or cell body

- Is where cell metabolism takes place
- Has places where messages from other neurons can be received (called a Post Synaptic Element)
- Contains many other structures related to metabolism
 - Mitochondria
 - Endoplasmic Reticulum
 - Golgi apparatus
 - Other structures
 - These structures are not important for this class.

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Axons and Related Structures

- Axons are structures that send information to other neurons or muscle cells.
- Have many structures

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Axon Hillock

- Is at the base of the axon
- Is the place where neurons decide to send a signal (called an action potential) to another neuron

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Body of the Axon

- This structure can branch (called a collateral)
- Branching continues into smaller and smaller branches called Teleodendria

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Presynaptic Element

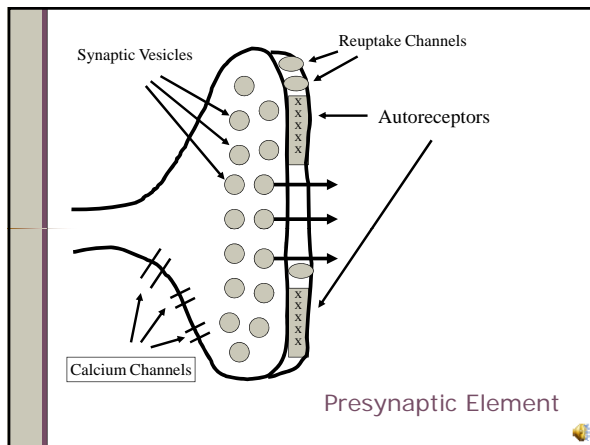
- Also called terminal buttons, terminal boutons, and other names)
- We will call it presynaptic element

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Contains Several Structures

- Synaptic Vesicles (sacks)
 - Sacks contain chemicals called neurotransmitters
- Presynaptic Membrane
 - Autoreceptors
 - Reuptake channels
- Receptors from other neurons
- Calcium Channels

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Axons can be one of two types

- Myelinated
 - Myelin is a fatty covering over the axon
 - Helps to increase the speed of the action potential
 - The more myelin there is, the faster the speed of the action potential

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Non-Myelinated axons

- Many axons do not have myelin
- Are slower than myelinated axons
- However, the fatter the axon is, the faster the action potential will go.

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Dendrites

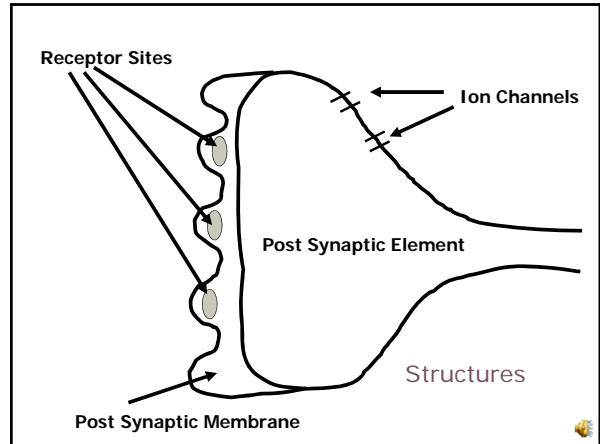
- Some neurons do not contain this structure.
 - only have soma's and axons
- Generally only receive information
- Contain a post synaptic element
 - Has a post synaptic membrane
 - Have receptor sites to receive neurotransmitters

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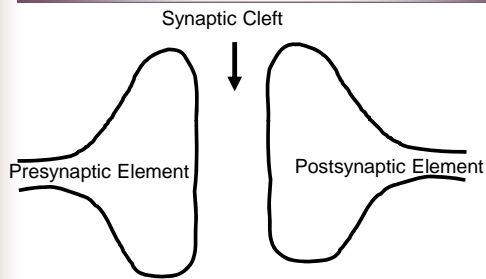
Dendrites

- So, both Dendrites and Soma's can receive information. Both contain a post synaptic element.

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Space



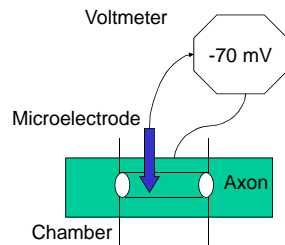
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How Neurons Work

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Measuring the Neuron Resting Membrane Potential

- Giant axon from a squid is placed in seawater in a recording chamber.
- Glass microelectrode is inserted into axon.
- Voltage measures -70 mV inside with respect to outside



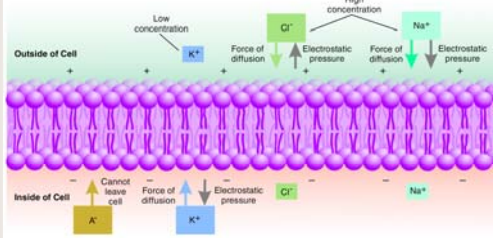
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How Neurons Work

- Based on concentration gradients of four ions
- Sodium (Na), Potassium (K), Chloride (Cl), and Structures inside the axon called Anions (A)
- Sodium and Potassium are positively charged and are balanced out by chloride and anions

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Membrane Ion Concentrations



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How Neurons Work Cont.

- Normally some sodium leaks into the axon.
- But cells don't like sodium, so they have pumps that remove sodium called sodium potassium pumps.
- The pumps remove sodium to the outside.

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How Neurons Work Cont.

- The inside of axons have lots of potassium and anions and are negatively charged. Potassium also leaks to the outside of the neuron
- The outside of axons have lots of sodium and chloride and are positively charged.
- When an axon is at rest, the outside of the axon is positively charged and the inside is negatively charged.
 - Called the Resting Potential

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Concept

- **Influx** Material (Ions) moving to the inside of a membrane.
- **Efflux** Material (Ions) moving from the inside to the outside of a membrane.
- **Equilibrium** Where material (ions, concentrations, etc.) are equal on both sides of a membrane.

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Why do Ion's Move?

- **Concentration Differences**
 - Compounds move from high concentrations to lower concentrations.
- **Electrostatic Pressure**
 - Like charges repel each other
 - Opposite charges attract each other.

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Why is There a Resting State?

- The neuron membrane is selectively permeable to certain ions.
- Sodium Na^+
- Potassium K^+
- Chloride Cl^-
- Calcium Ca^{++}

- When Na is at equilibrium (+55mV) there is a large efflux of K
- When K is at equilibrium (-70mV) there is a influx of Na
- Nature thus splits the difference. (approx. -65mV)
 - Get some efflux of K
 - Get some influx of Na
- Best solution uses the least amount of energy (ATP)

Result

- Balance between the ions moving through the membrane makes the resting state
- Note: the resting state can range between -30mV to -65mV

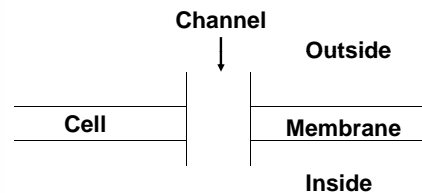
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Axon Pores

- Axons have two types of channels or pores for different ions.
 - Passive Channels
 - Are open all of the time and allow ions to pass through the membrane.
 - Active or Gated Channels
 - Are activated by changes in voltage
 - E.g., voltage-gated Na channels

Passive

Are open all the time



- Many K passive channels are open
 - Some Na passive channels are also open
 - Channels are selective
 - Na channels are only selective for Na, K doesn't get through.
 - Ions go through channels at different rates.
 - 12 K to 1 Na
- Some ions are moving but not many. It lots of movement you get equilibrium.

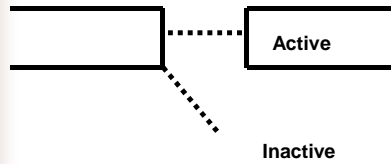
Voltage-Gated Channels

- Are needed for the action Potential.
- Membrane potential must depolarize 15mV before voltage-gated channels open.
- Thus, only certain receptors sites close to the hillock will open voltage-gated channels.

Voltage-Gated Channels

- Have two gates
 - Active - Waiting for thresholds
 - Inactive - Waiting for positive charges or zero point

Sodium Voltage Gated Channel



Active Gates are Usually Closed
With Depolarization, Active Gates Open

Steps

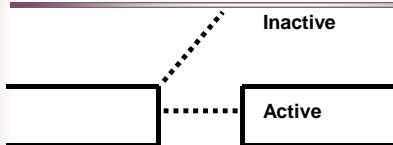
- Get Depolarization
- If depolarize 15mV – Active Gates Open
 - Get an action potential
- If does not depolarize 15mV – Nothing

So, All or nothing
Is happening at the Axon Hillock.

Characteristics

- Several types – Na, K, Ca
- Channels are only one way.
- Channels are Ion specific
- Active gates open at depolarization, then inactive gate closes at zero or above
- After inactive gate closes, Active Gate closes, inactive gate then opens.
- Repeat.

Potassium Voltage Gated Channel



Active Gates are Usually Closed
With Depolarization, Active Gates open

Points

- Like the Na Channel, the gates are sluggish
- It takes awhile for them to close

Summary

1. **Active Channels are usually closed. They open when you depolarize them.**
2. **Once Na has entered and begins to approach equilibrium, the inactive gate begins to close and ultimately stops Na from entering**
3. **The inactive gate is sluggish – thus, it takes awhile to close. Consequence, you get a positive overshoot in the process of the action potential.**

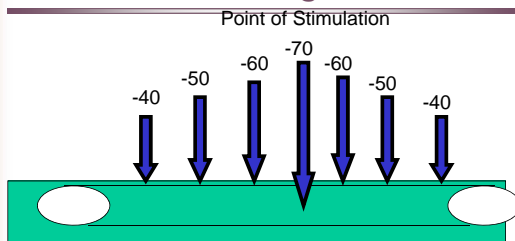
4. When Na enters, the pumps start up and begin to remove Na and bring in K (3Na/2K)
5. Finally inactive gate closes.
6. After enough Na is removed and K enters, the active gate closes and inactive gate opens
7. Process repeats again.

Types of Potentials

Electrotonic Potentials also called Passive Potentials

- Are decremental – they decay
 - Rate of decay depends on
 - Amount of Myelin
 - Length of the axon.
 - Diameter of the axon

Decremental Changes



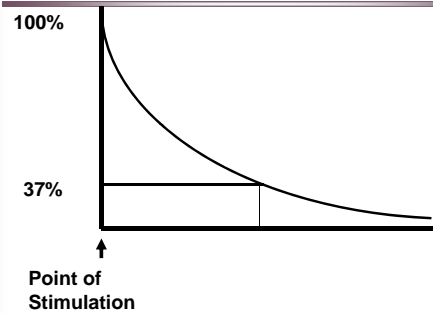
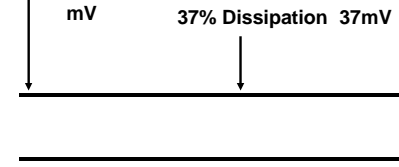
Charges Decrease in Both Directions

- Begin to stimulate, get depolarization (becomes more positive)
- K begins to move against the membrane and tries to leave by passive diffusion.
- If the membrane is thick, the length of dissipation also increases.

Length Constant

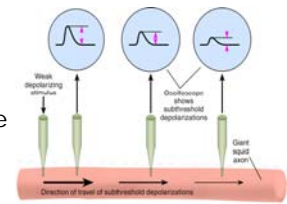
- Distance along the membrane from the point of stimulation where the change in the resting potential has dissipated to 37% of the original

Point of Stimulation 100 mV



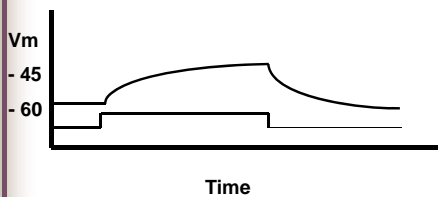
Summary

- Local potentials degrade with time and distance
- Local potentials can summate to produce an Action Potential



Temporal Summation

- Present a stimulus every ____ milliseconds.
- Observe depolarization



Resistance

- If membrane is resistant to K leaving, it becomes more difficult or K to leave
- Speed is the same
- All is done by passive channels.
- Decrease resistance by increasing the diameter of the axon.

Thus, length constant will also dictate temporal summation

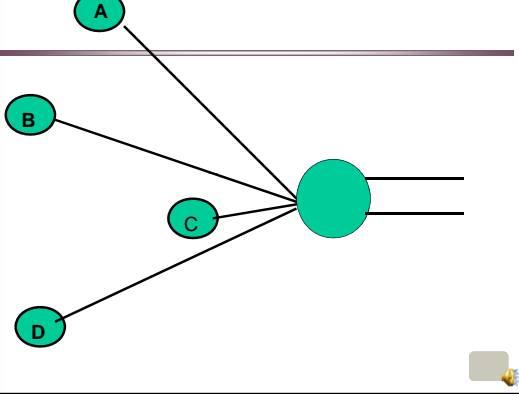
Action Potential

- Occurs because voltage-gated channels open
 - (Different from Passive Channels)
- Results in rapid and large Na influx
- The inside of the neuron becomes more positive (depolarization) and then becomes more negative (hyperpolarization)

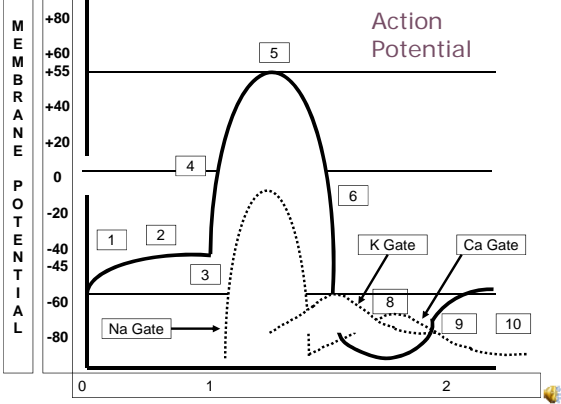
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Influences

- Distance away - Length constant
- Temporal summation
- Inhibitory neurons
 - Cause hyperpolarization to the hillock (becomes more negative)
- Combination of depolarization and hyperpolarization determines if Voltage-Gated channels open.



- C alone Voltage channels open – get AP
- A alone Get Nothing
- B alone Get Nothing
- D alone Get Nothing
- A + B + D Voltage channels open – get AP



Process of an Action Potential

1. Stimulation Begins
2. K begins to leave by passive channels. Na enters by passive channels.
 - Get a change in concentration gradients
 - Amount that leaves depends on the strength of the stimulus, how often it occurs, etc.
 - Begins to become more positive (depolarization).
3. If depolarization is reaches 15mV, voltage-gated Na channels open.
 - Sodium enters the neuron (influx)
4. Sodium-Potassium Pumps (Na K ATPases) start
 - Removes Na and brings in K
 - K also leaves through passive channels.
 - K voltage gated channel begins to open about ½ millisecond after Na voltage gated channels open

5. Na voltage-gated channel finally closes,
 - Action potential begins to fall
6. K gate finally closes
 - K is still leaving by passive channels
 - Na is leaving by Na K pump
 - Action potential continues to fall
 - Get a negative undershoot from resting state.
7. Ca voltage-gated channel begins to open
8. Still do not have enough K so Calcium continues to enter the neuron (influx)
9. Finally enough K has entered so the Calcium channel begins to close
10. Ca channel finally closes
11. Process Repeats

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Review

- When a stimulus enters a receptor on a dendrite, it causes change in polarity.
- Causes a change in the chemical concentration gradients.
- Allows sodium to enter in small amounts and depolarizes (makes it more positive) the neuron.
- The depolarization travels to the axon hillock. If the charge depolarizes the hillock 15mv, get an action potential.
- If the charge is not strong enough, the signal stops.

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Review cont.

- Causes sodium gates in the axon to open.
- Get Na Influx
- Result, the axon goes from negative on the inside to positive on the inside.
- This change goes down the axon like a wave.
- After the sodium enters, the sodium potassium pumps turn on and begin removing sodium.
- Also goes down like a wave

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Review

- So we have two waves going down the axon,
 - The sodium entering the axon
 - The sodium being pumped out
- Ultimately the result is a negative undershoot

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When the axon potential reaches the presynaptic element

1. It causes calcium (Ca) to enter the presynaptic element.
 - Calcium causes the synaptic vesicles to bind with the presynaptic membrane
 - The neurotransmitter is then released into the synaptic cleft.
 - The neurotransmitter crosses the cleft and binds on receptors in the post synaptic element on either the dendrite or soma.
2. Causes a small electrical charge and the process repeats itself.

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How neurotransmitters (NT) are removed from receptors

- NT is removed two ways:
 1. It is degraded by enzymes made by glial cells or within the post synaptic membrane
 2. It is reabsorbed into the presynaptic element.

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Impact of Drugs

- Impacts the neuron several ways
 - Entire neuron (Alcohol)
 - Presynaptic Element (Cocaine, Meth.)
 - Postsynaptic Element (Opiates)
 - Specific receptor sites (Barbiturates)
 - All of the above (Alcohol).

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Entire Neuron

- Alters the lipid bilayer of the neuron
- Slows ion flow
- Ultimately reduces the height of the action potential
- Ultimately reduces Calcium influx
- Fewer NT is released
- Less stimulation on post synaptic element
- Less depolarization in the next neuron
 - Does not become as positive
- Result - Fewer action potentials in the next neuron

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Presynaptic elements

- Drugs block the reabsorption of the NT
- Result, NT remains on post synaptic receptors longer
- Get more action potentials

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Post Synaptic Elements

- Blocks the NT from binding on the receptor
- Less depolarization
- Fewer action potentials
- Depending on the brain area impacted (medulla) can cause death or temporary memory loss (hippocampus).

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Conclusions

- Very important area
- Has had tremendous impacts in our understanding of drug effects (positive and negative)

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