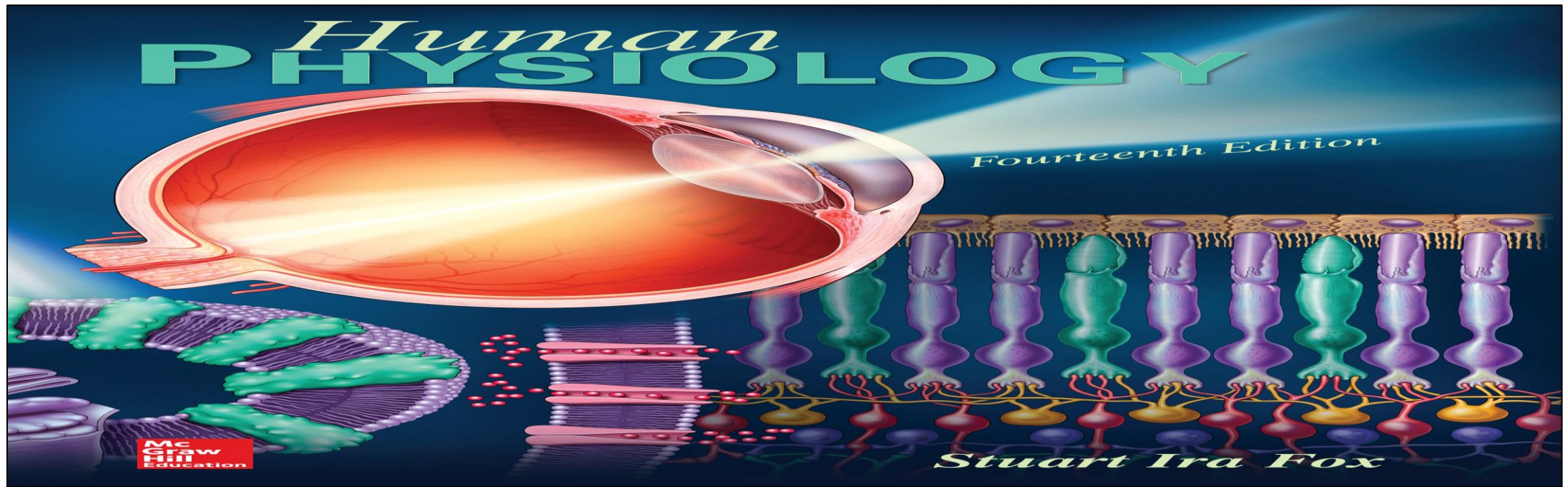


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د. محمد الكفاوين



# Physiology for Nursing

## Dr. Mohammad Al-Kafaween

Text Book: HUMAN PHYSIOLOGY, FOURTEENTH EDITION  
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# What is Anatomy? What is Physiology?

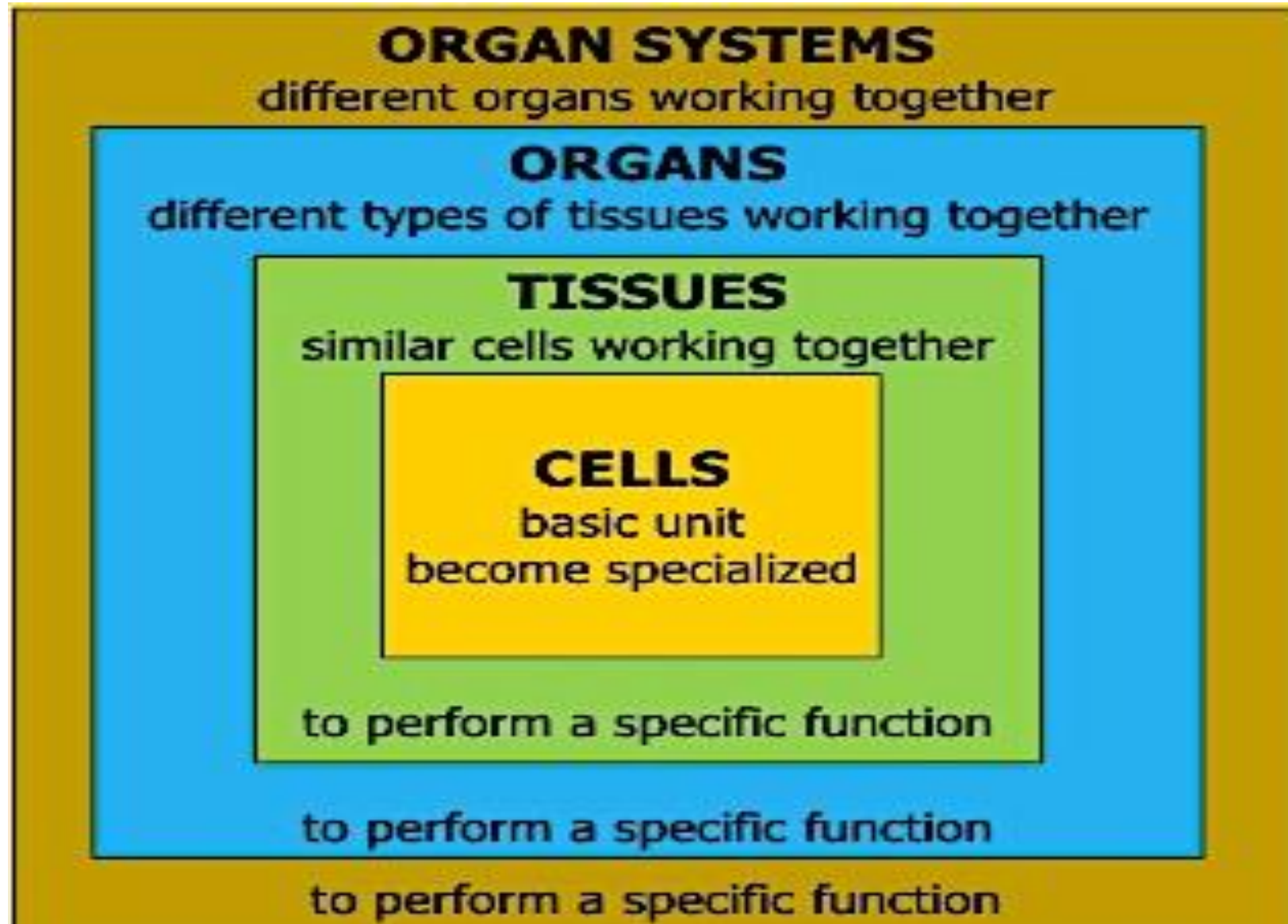
- Anatomy and Physiology are closely related.
- **Anatomy** – study of the structure and shape of the body and its parts (and how those structures relate to each other).
  - Ex: Parts of a heart
- **Physiology** – study of how the body and its parts work or function
  - Ex: How the heart pumps blood
- Physiology is explainable only in terms of the underlying anatomy.

# What is Physiology?

- Study of biological function; how the body works.
- Concerned with the normal function
- Emphasizes mechanisms
- Explained by cause-and-effect sequences
- Based on scientific experiments



# What is the role of human physiology in pharmacy?



# Chapter 1: Homeostasis

- Homeostasis is constancy of the internal environment.
- The main purpose of our physiological mechanisms is to maintain homeostasis.
- Deviation from homeostasis indicates disease.
- Homeostasis is accomplished most often by negative feedback loops.

# Negative Feedback Loops

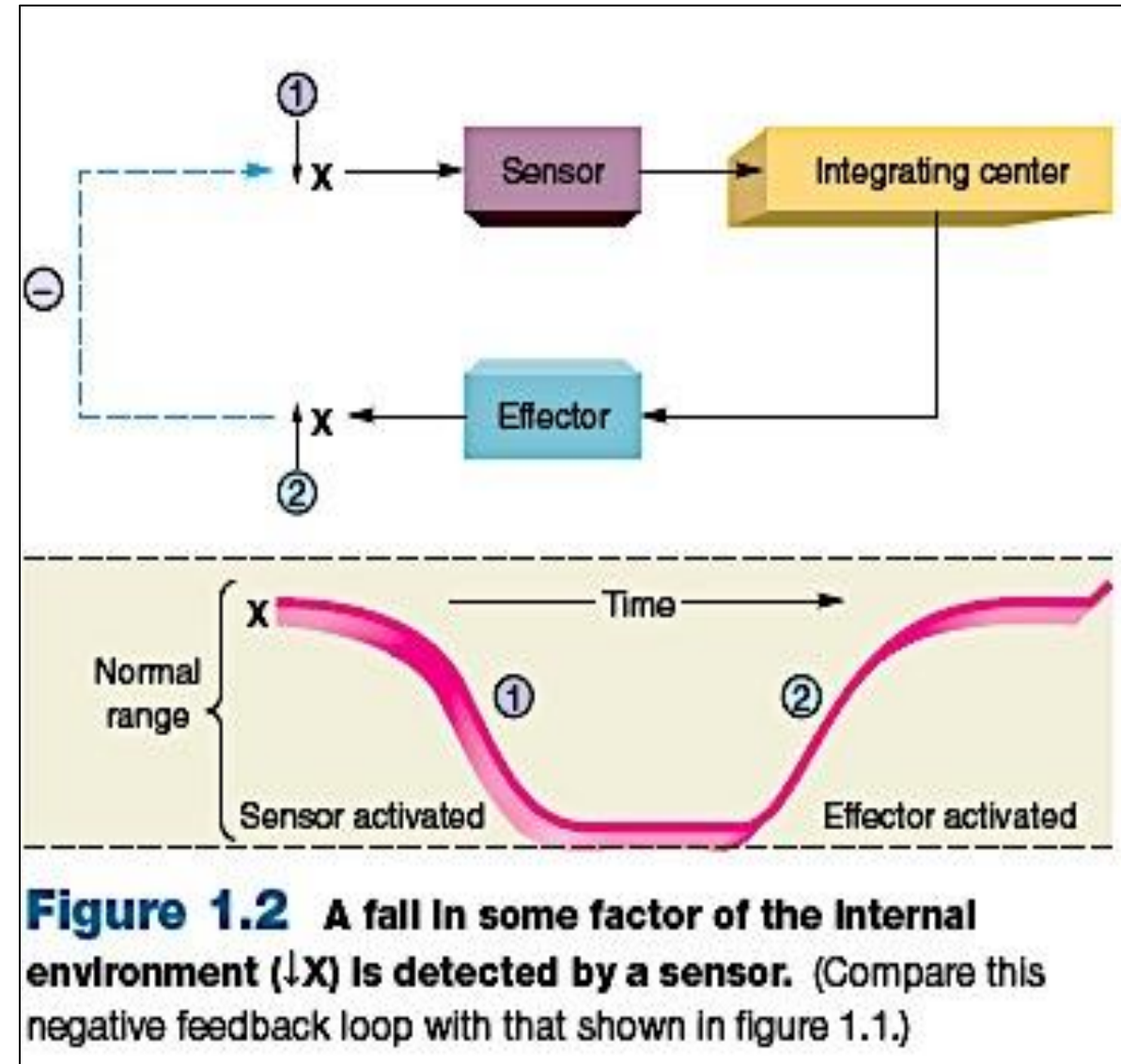
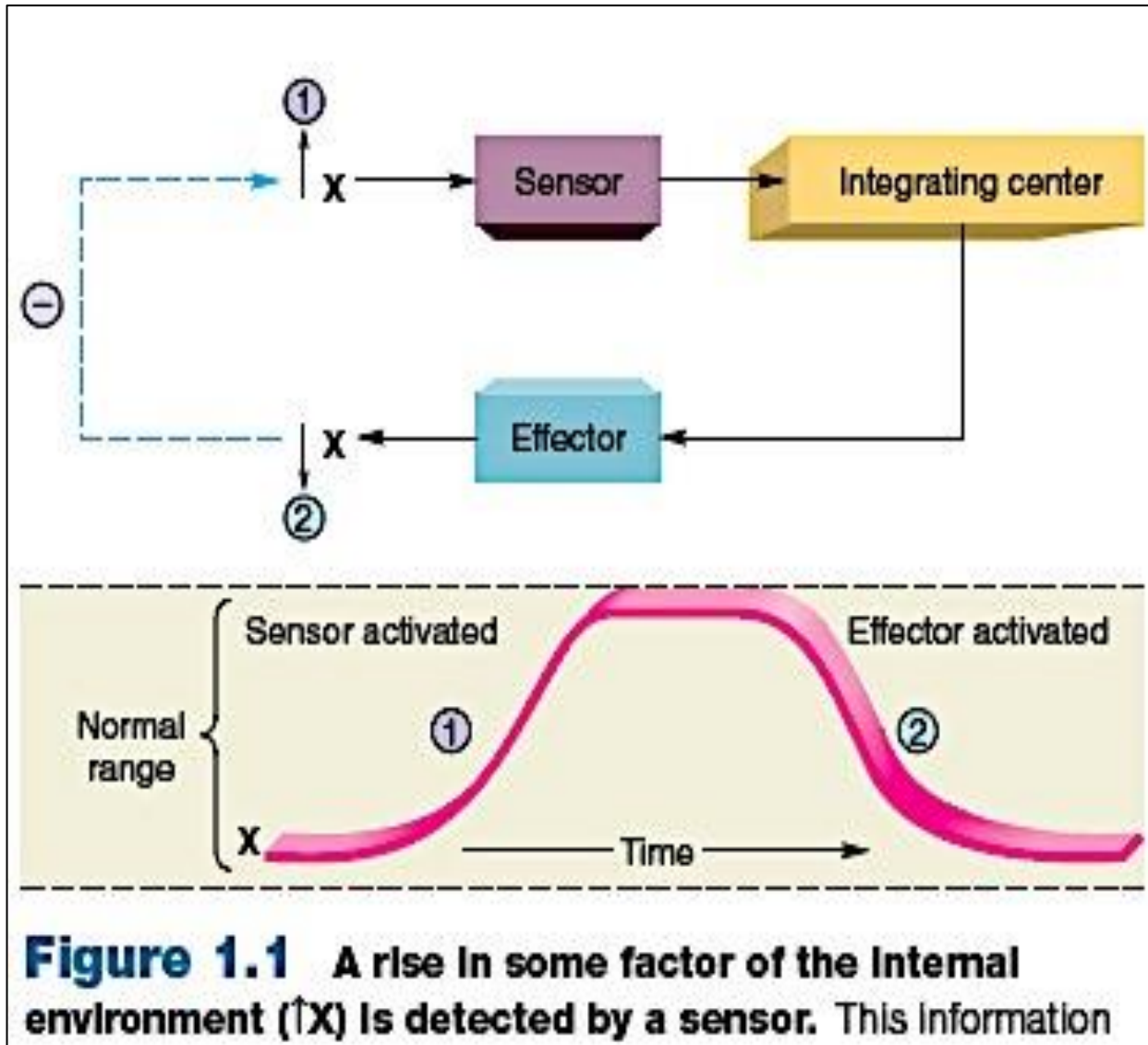
- Pathway

**A. Sensors** in the body to detect change and send information to the:

**B. Integrating center (Brain, CNS, endocrine glands)**, which assesses change around a **set point** (house thermostat). The integrating center then sends instructions to an:

**C. Effector (muscle or gland)**, which can make the appropriate adjustments (**increase or decrease**) to counter the change from the set-point

# Negative Feedback Loops

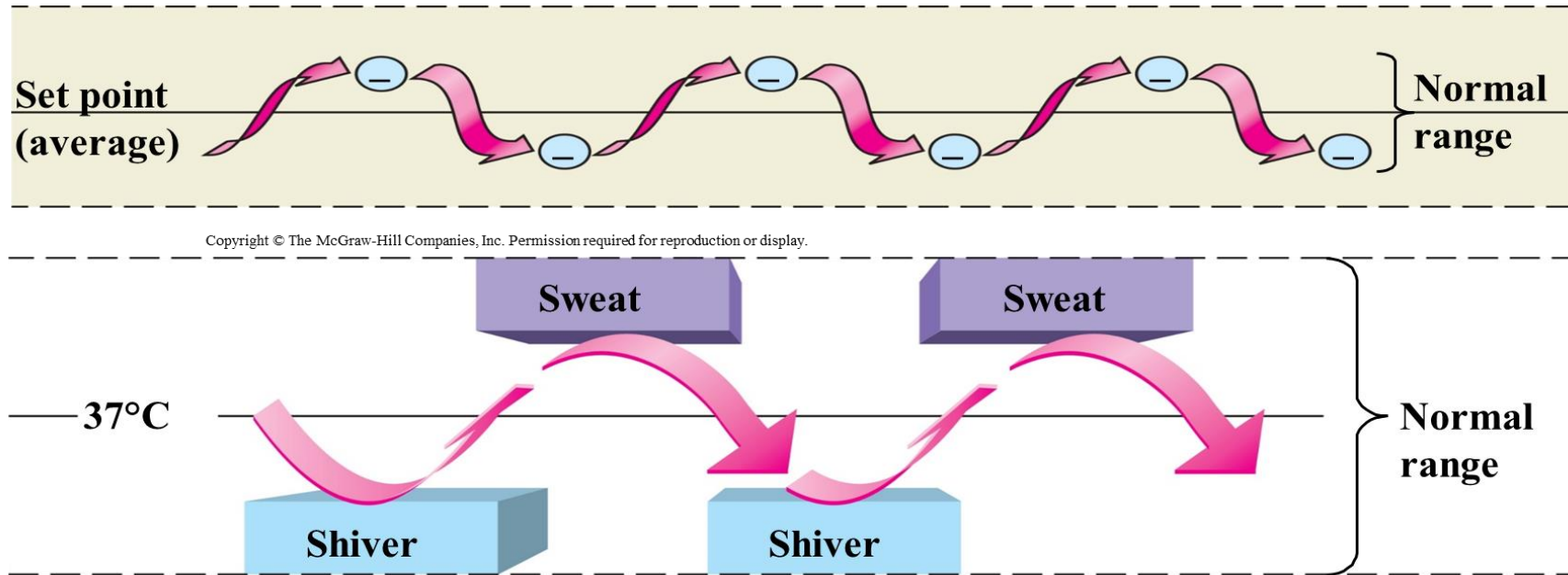


## The regulation of body temperature (Example of negative feedback loops)

- a. Sensors in the brain detect deviation from 37° C. Another part of the brain assesses this as actionable, and effectors (sweat glands) are stimulated to cool the body.
- a. Once the body is cool, sensors alert the integrating center, and sweat glands are inhibited.
- a. The end result regulates the entire process. Production of the end product shuts off or down-regulates the process. Therefore, it is called a negative feedback loop.

# Antagonistic Effectors

- Homeostasis is often maintained by opposing effectors that move conditions in opposite directions.
  - 1) This maintains conditions within a certain normal range, or dynamic constancy.
  - 2) When you are hot, you sweat; when you are cold, you shiver. These are antagonistic reactions.

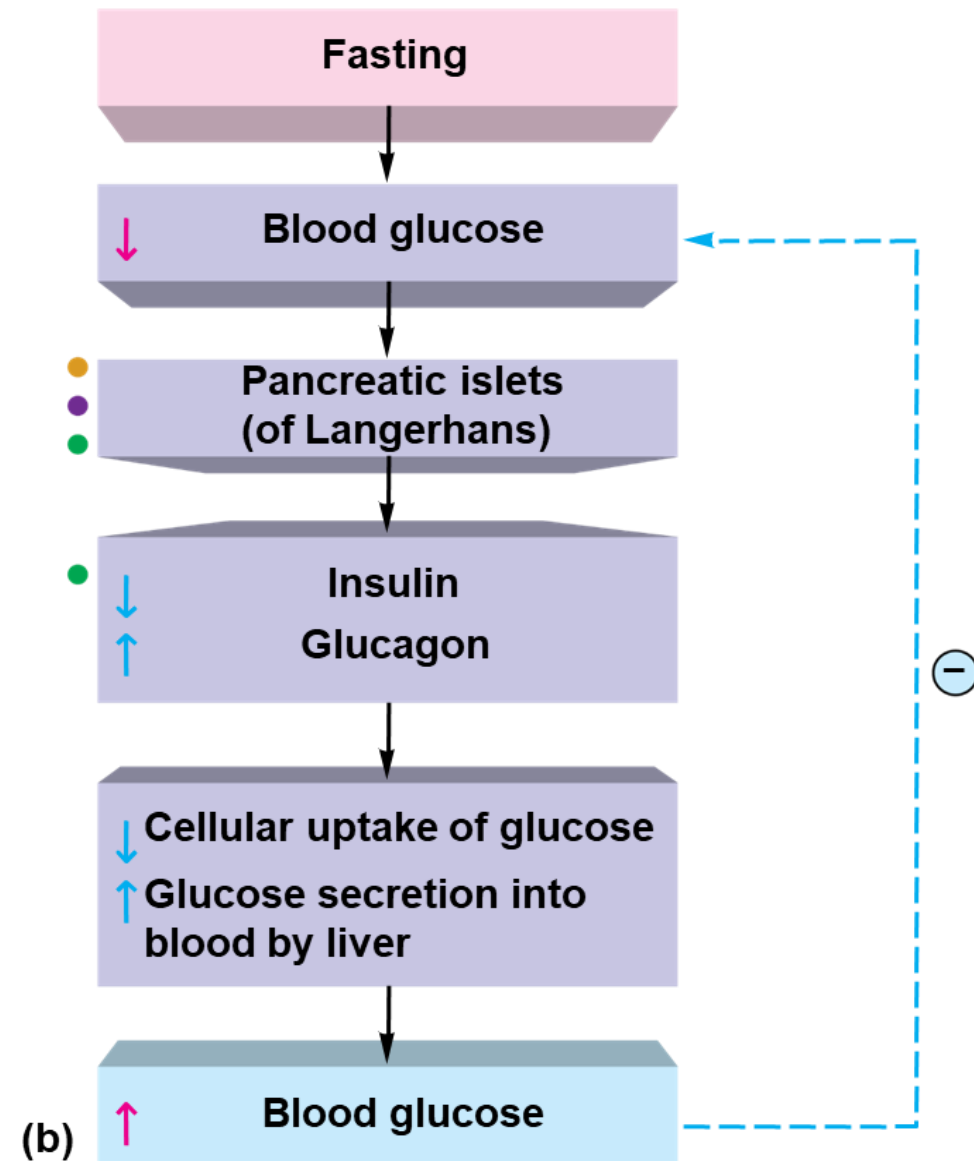
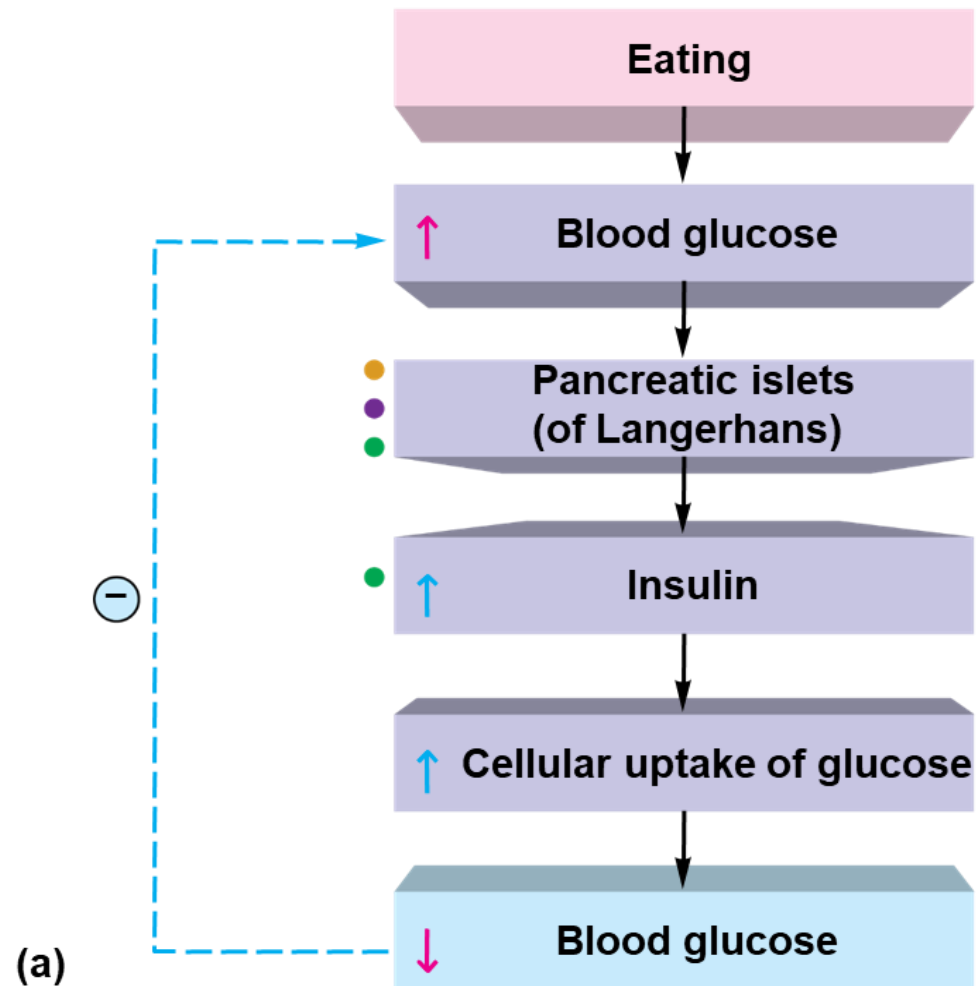


Negative feedback loops are continuous



# Blood glucose levels: an example of antagonistic effectors

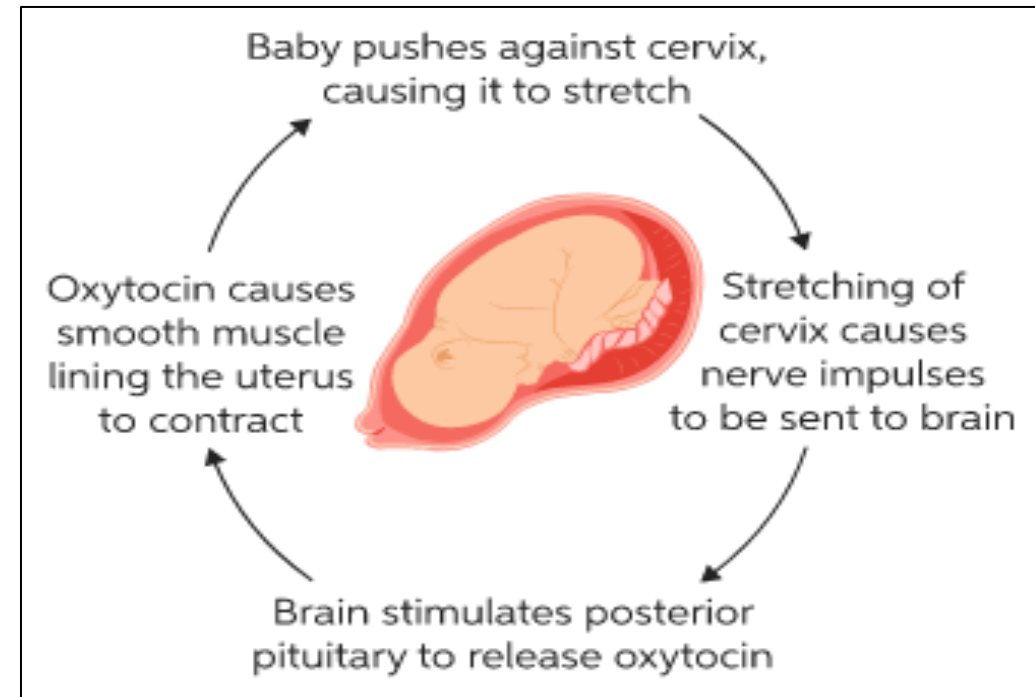
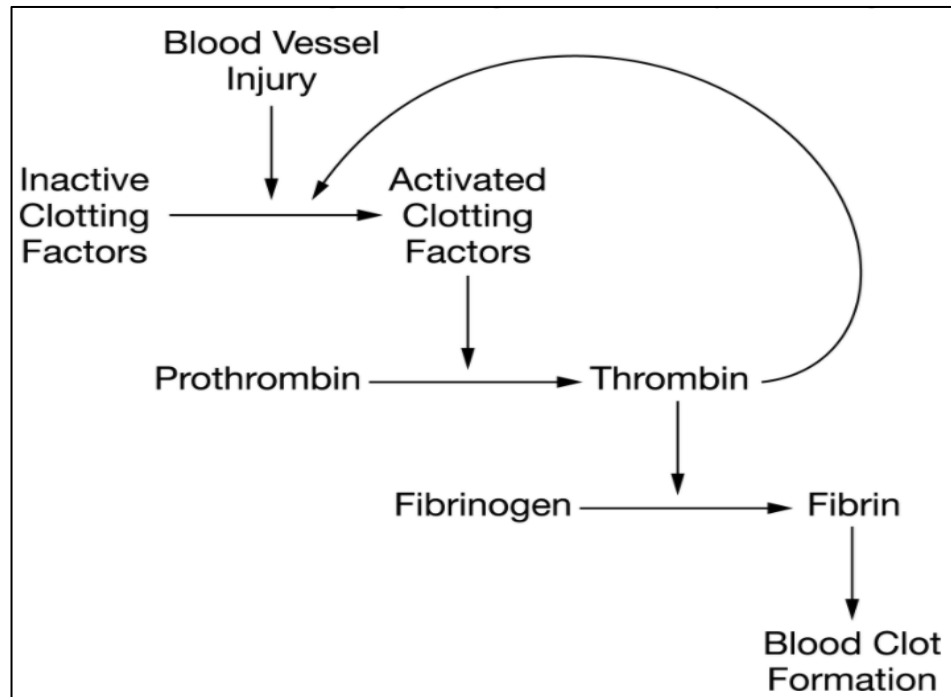
- Sensor
- Integrating center
- Effector





# Positive Feedback

1. The end product in a process stimulates the process.
2. The action of effectors amplifies the changes that stimulated the effectors
3. Positive feedback could not work alone, but it does contribute to many negative feedback loops.
  - a. For example, if a blood vessel is damaged, a process is begun to form a clot. Once the damage is fixed, clotting ends (negative feedback). However, the process of forming the clot involves positive feedback.
  - b. The strength of uterine contractions during childbirth is also regulated by a positive **feedback loop**.



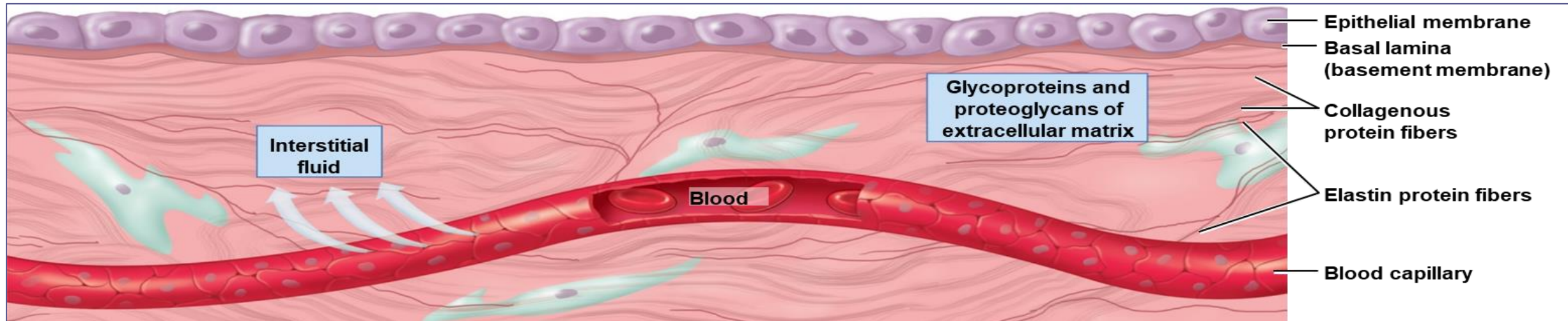
# Intrinsic and Extrinsic Regulation

- Regulation of processes within organs can occur in two ways:
  - a. **Intrinsically**: Cells within the organ sense a change and signal to neighboring cells to respond appropriately.
  - a. **Extrinsically**: The brain (or other organs) **regulates** an organ using the **endocrine or nervous system**.
- The **nervous system** “innervates” organs with nerve fibers.
- The **endocrine system** releases hormones into the blood, which transports them to multiple target organs.

# **Chapter 6: Interaction between cells and extracellular environment**

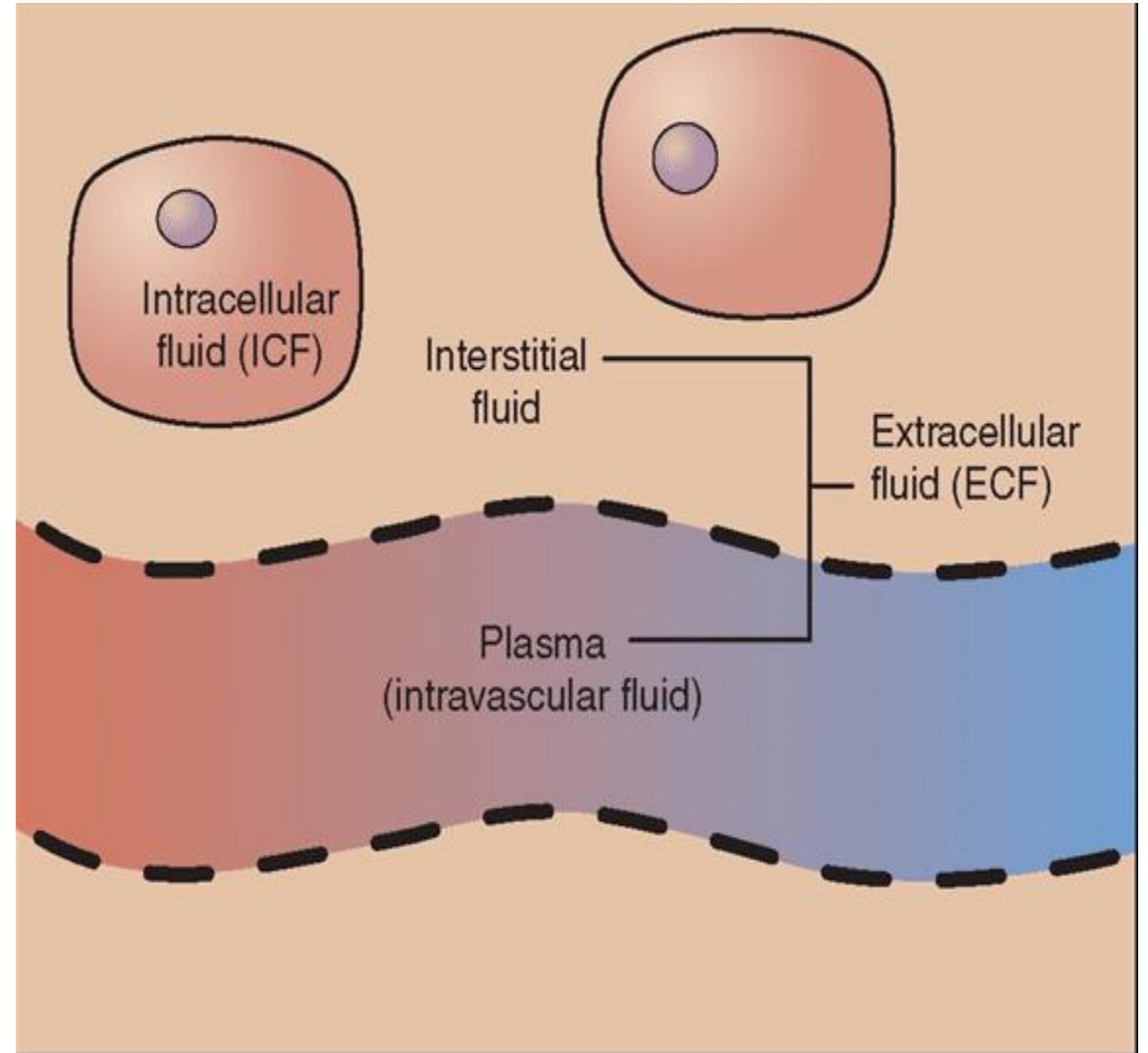
# Chapter 6: Extracellular Environment

1. The extracellular environment includes everything located outside the cells.  
Extracellular Matrix: contains protein fibers of collagen and elastin, and a gel-like ground substance
2. Cells receive nourishment from and release wastes into the extracellular environment.
3. Cells communicate with each other by secreting chemical regulators into the extracellular environment.



# Body Fluids

- **67% of our water** is within cells in the **intracellular compartment**.
- **33% is in the extracellular compartment**. Of this:
  - ❖ 20% is in **blood plasma**.
  - ❖ 80% makes up what is called **tissue fluid, or interstitial fluid**; connects the intracellular compartment with the blood plasma.



# Plasma membrane transport

- Plasma membrane permeability:
  - a. The plasma membrane is **selectively permeable**, meaning that it allows **some molecules to cross but not others**.
  - b. Generally not permeable to proteins, nucleic acids, or other large molecules.
  - c. Generally permeable to ions, nutrients, and wastes

# Diffusion and Osmosis

1. **Solution**: consists of a **solvent** (water) and a **solute** (molecules dissolved in water)

a. Molecules in a solution are in a constant state of motion.

b. If there is a concentration difference between two regions, random motion will establish equilibrium via diffusion.

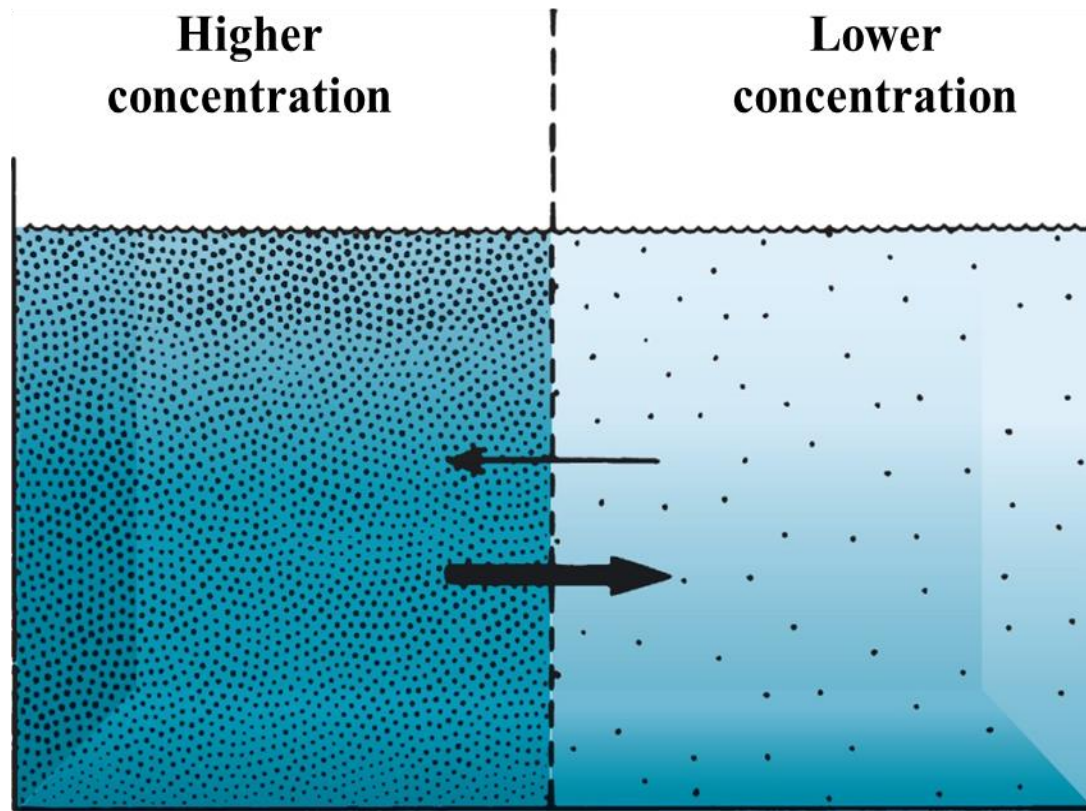
2. **Diffusion** will occur without a physical separation or across a permeable membrane

A. Net diffusion - due to random movement, the net direction of diffusion is from **high to low** solute concentration.



# Diffusion of a Solute

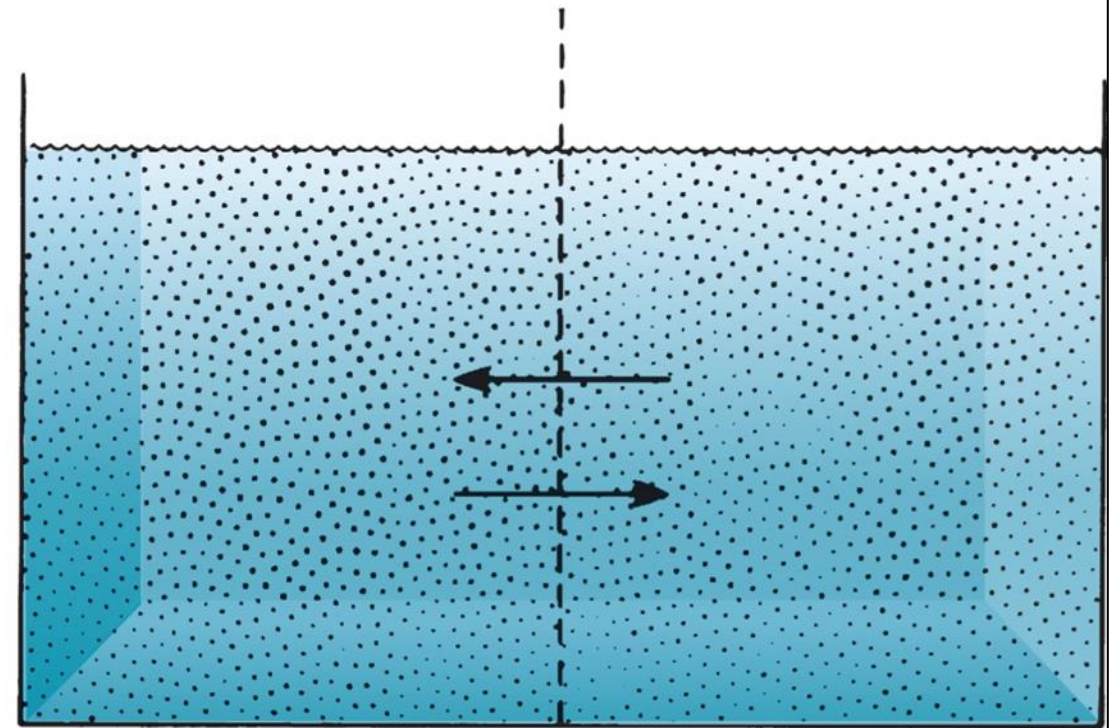
(a)



Net diffusion

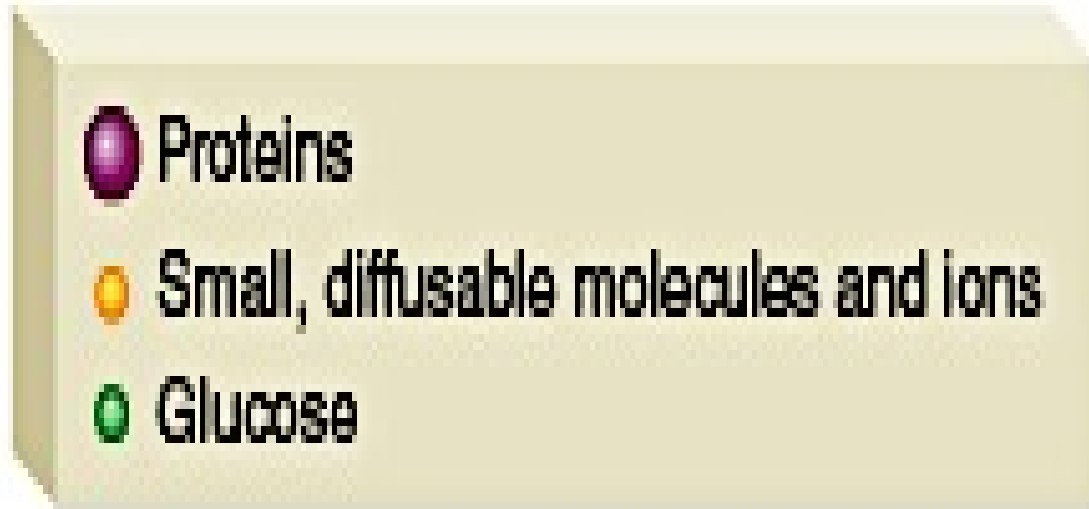
(b)

Equal concentrations

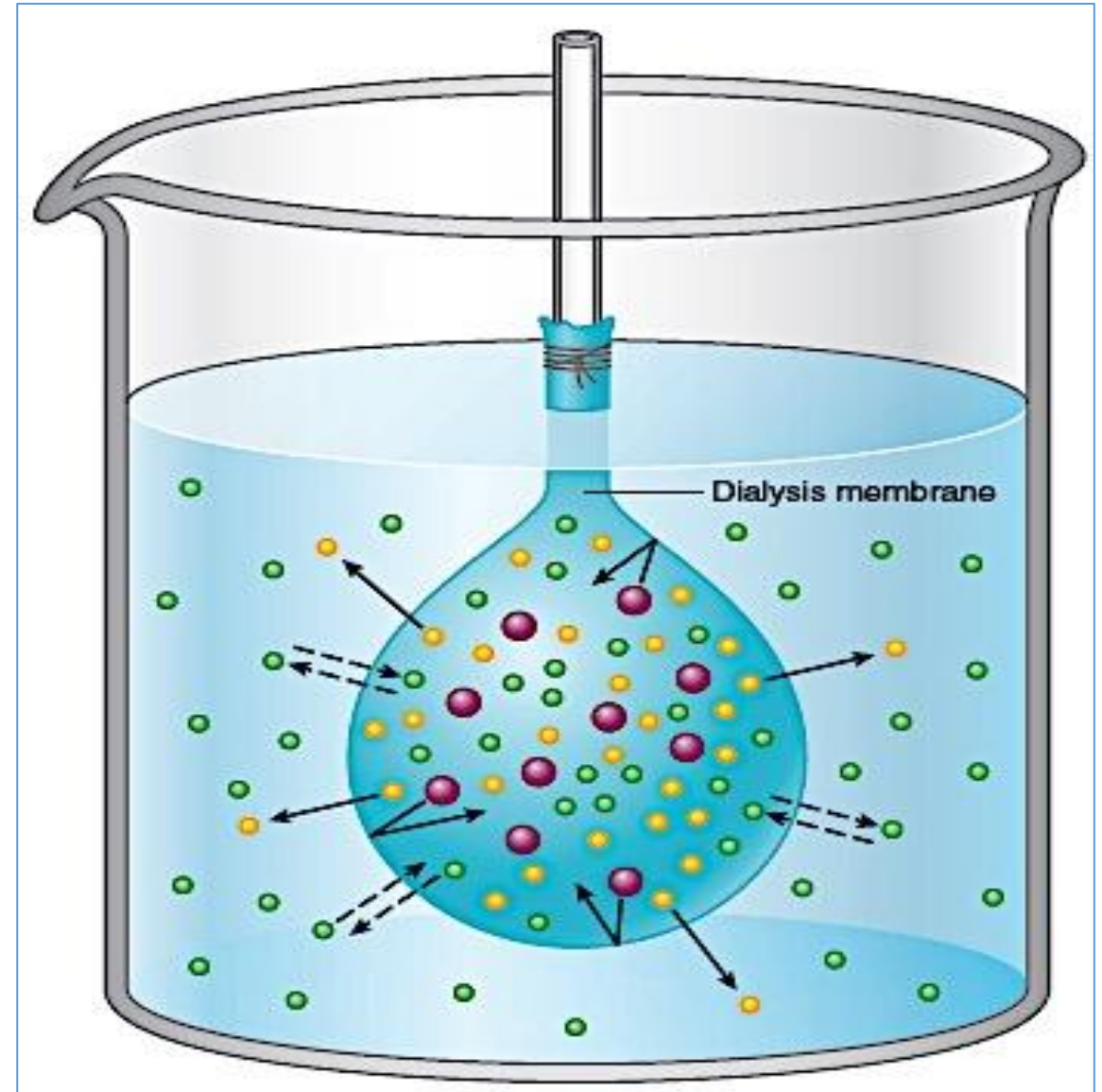


No net diffusion

# Diffusion Through a Dialysis membrane



Dialysis is the separation of smaller molecules from larger molecules in solution by selective diffusion through a semipermeable membrane.



# Diffusion through the Plasma Membrane

## Non carrier-mediated

**Simple diffusion**  
of lipid soluble  
molecules through  
the phospholipid  
bilayer  
ex. O<sub>2</sub>, CO<sub>2</sub>  
steroid hormones

**Simple diffusion**  
of ions through  
membrane channel  
proteins  
ex. Na<sup>+</sup>, Cl<sup>-</sup>

**Simple diffusion**  
of water molecules  
(osmosis) through  
aquaporin (water)  
channels

## Carrier-mediated

**Facilitated  
diffusion**  
needs carrier to  
pass through the  
plasma membrane

**Active transport**  
needs ATP  
against the  
concentration  
gradient  
through Pumps

passive transport

# Osmosis

- Osmosis is the net diffusion of water (the solvent) across the membrane.
- Water molecules do not carry a charge, so they can pass through the plasma membrane slowly.
- Aided by channels in the membrane called **Aquaporins** found in the kidneys, eyes, lungs, salivary glands, and the brain

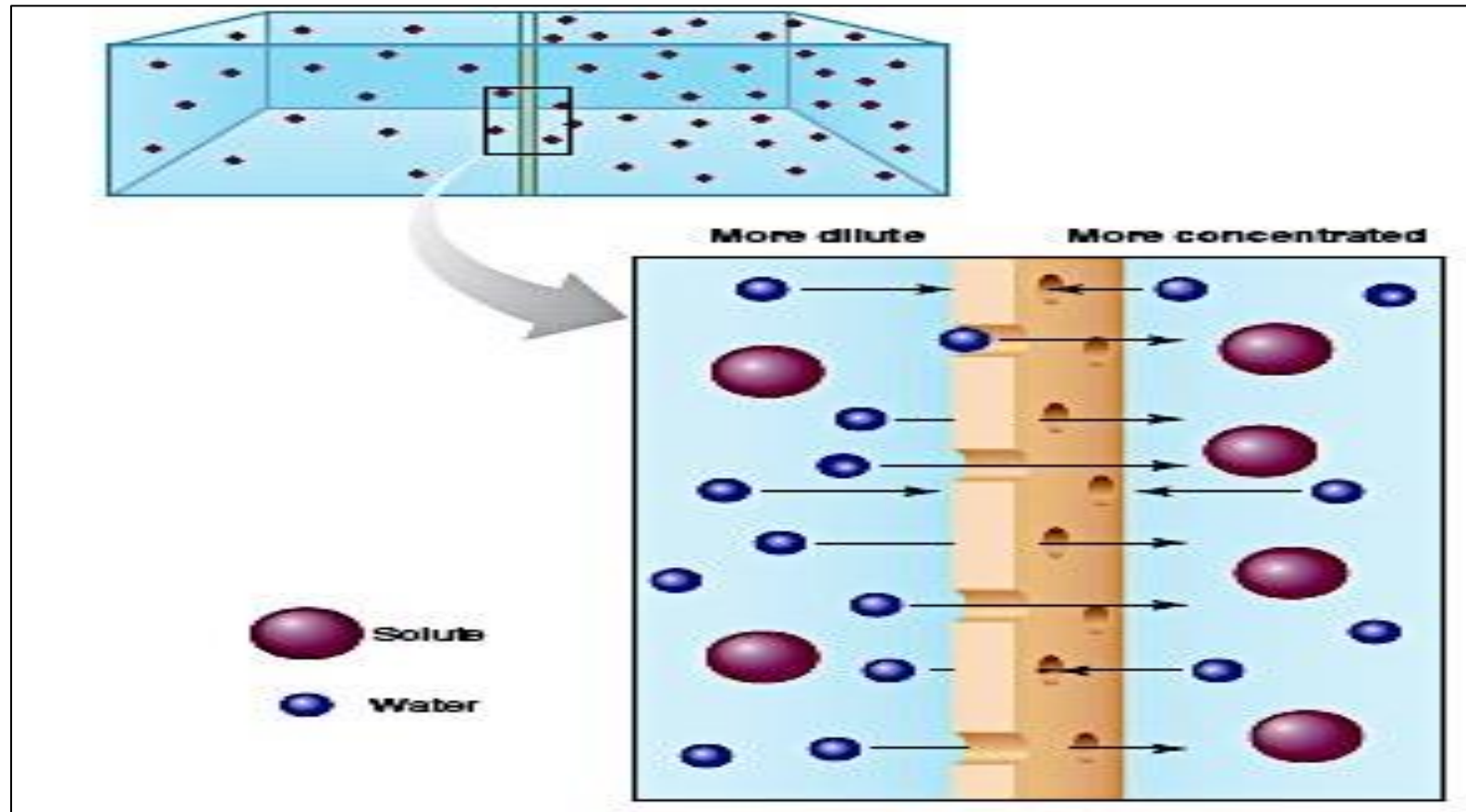
# Requirements of Osmosis

1. There must be a solute concentration difference on either side of a membrane permeable to water.
  2. The membrane must be impermeable to the solute
- Solutes that cannot freely pass through the membrane can promote the osmotic movement of water and are said to be **osmotically active**.



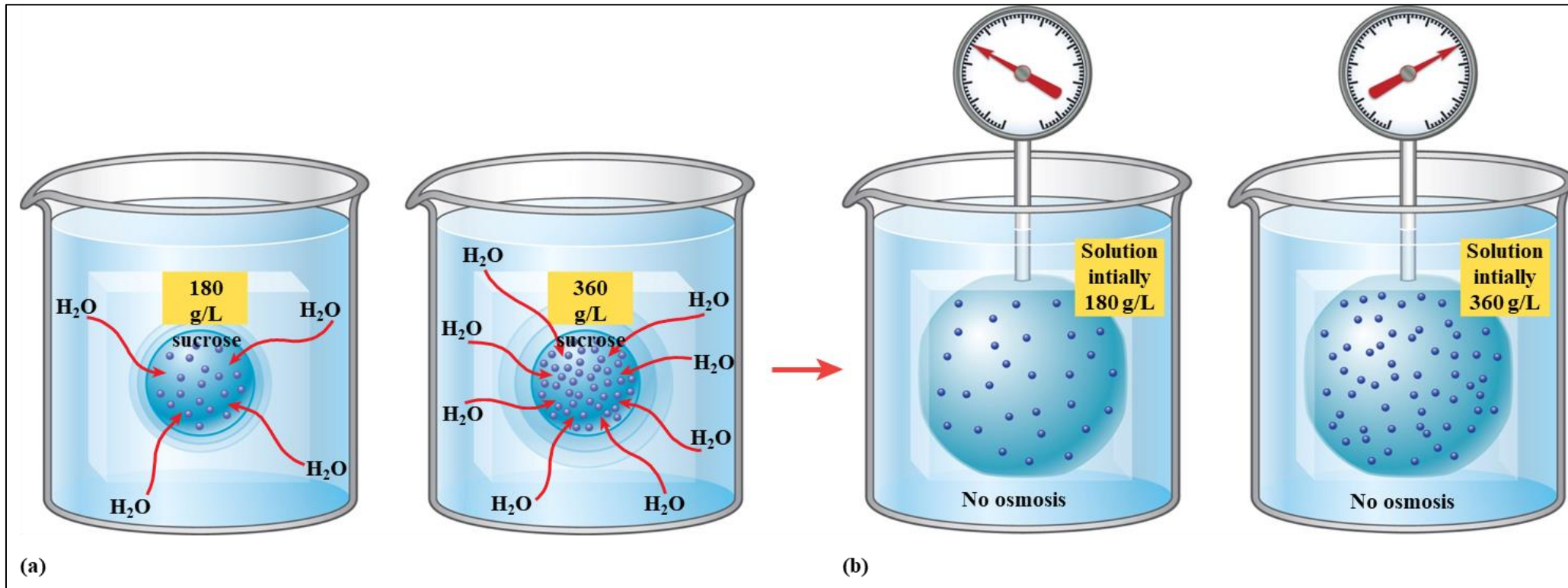
# Model of Osmosis

The net movement of water from the solution of lesser solute concentration (**higher water concentration**) to the solution of greater solute concentration (**lower water concentration**)



# Osmotic Pressure

- **Osmotic pressure** is the pressure needed to stop osmosis.
- It can be used to describe the osmotic pull of a solution. A higher solute concentration would require a higher osmotic pressure.
- Pure water has an osmotic pressure of zero



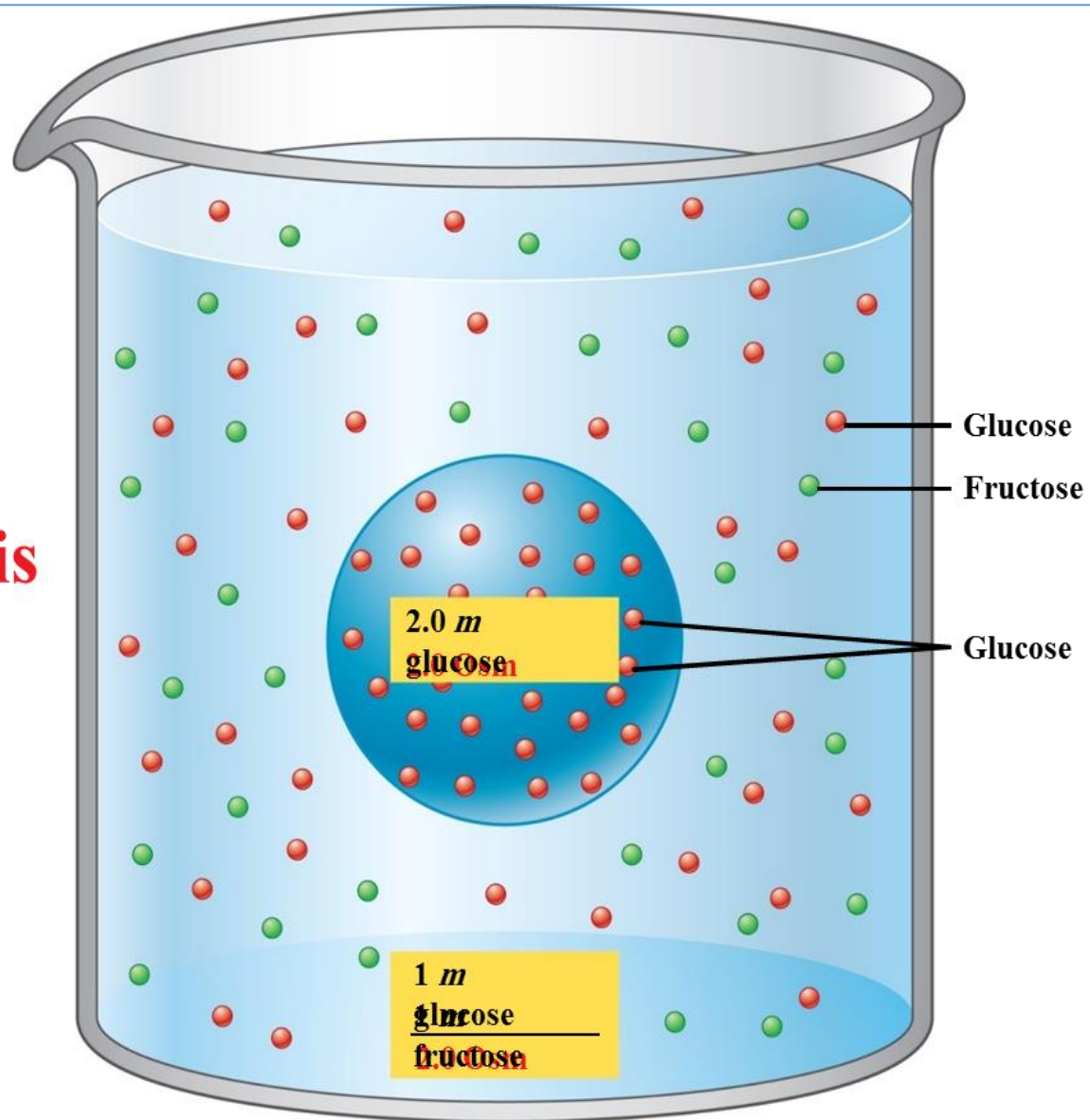


# Osmolality (Osm)

- Osmolality is the total molality of a solution when you combine all of the molecules within it.
- **Molality = moles solute / kilogram of solvent**
- Example: A (360 g (2m) glucose solution) has the **same osmolality** as a solution of (180 g glucose (1m) + 180 g fructose (1m)).
- These are both 2 Osm solutions.
- Electrolytes that dissociate in water have to be assessed differently.
  - NaCl dissociates into  $\text{Na}^+$  and  $\text{Cl}^-$  in water and must be counted as separate particles.
    - $\text{NaCl} \longrightarrow \text{Na}^+_{(\text{aq})} + \text{Cl}^-_{(\text{aq})}$
  - A 1m NaCl solution would actually be a 2 Osm solution.
- Osmolality can be measured by freezing point depression – how much the freezing point is lowered depends on the number of particles present in the solution

# Osmolality of sugar solutions

**Isotonic: no osmosis**

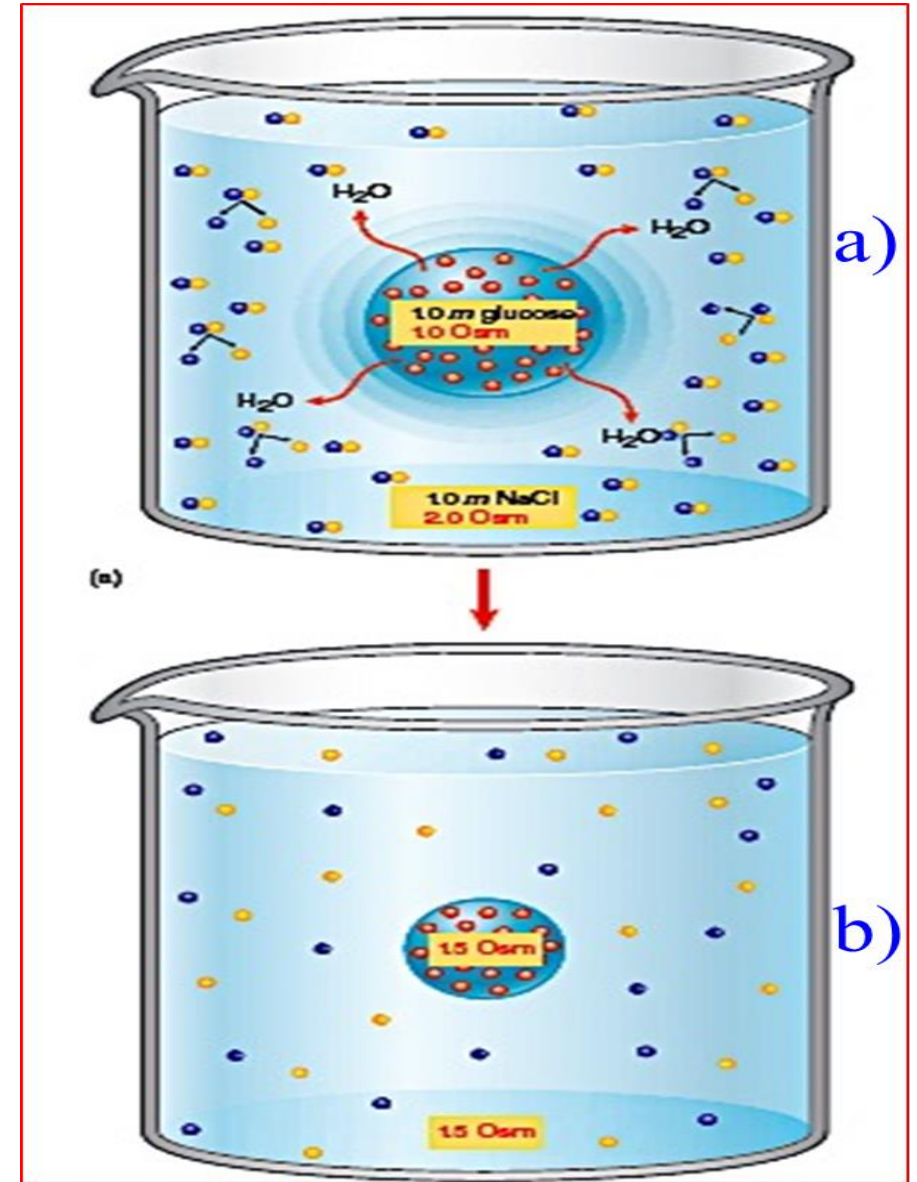


# Effect of Ionization on Osmotic Pressure

a) **Semipermeable membrane**: permeable to water but not to glucose, Na, or Cl

- water will move by osmosis into the NaCl solution, which has 2Osm concentration

b) **After osmosis**, the total concentration, or osmolality, of the two solutions is equal.

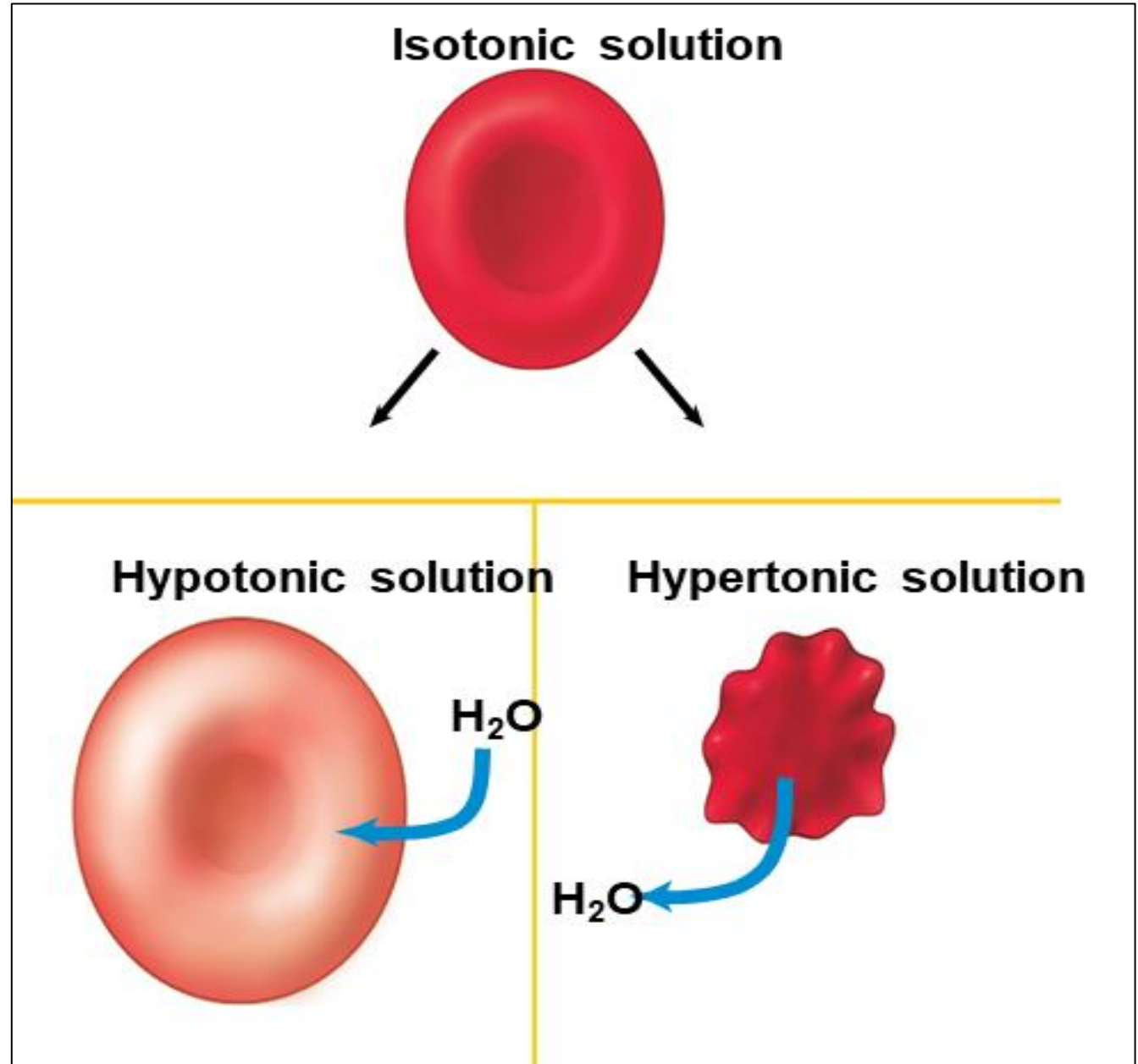


# Tonicity

- a. Plasma has the same osmolality as **a 0.3m glucose** or a **0.15m NaCl** solution.
  - 1) These solutions are considered isosmotic to plasma.
  - 2) Made as 0.9g NaCl/100mL water – normal saline
  - 3) 5% dextrose – 5g glucose/100 mL water
- b. **Tonicity** is the effect of a solute concentration on the osmosis of water.
  - 1) If a membrane separates a 0.3m glucose solution and a 0.15m NaCl solution, there will be no net movement of water = isotonic.

# Tonicity

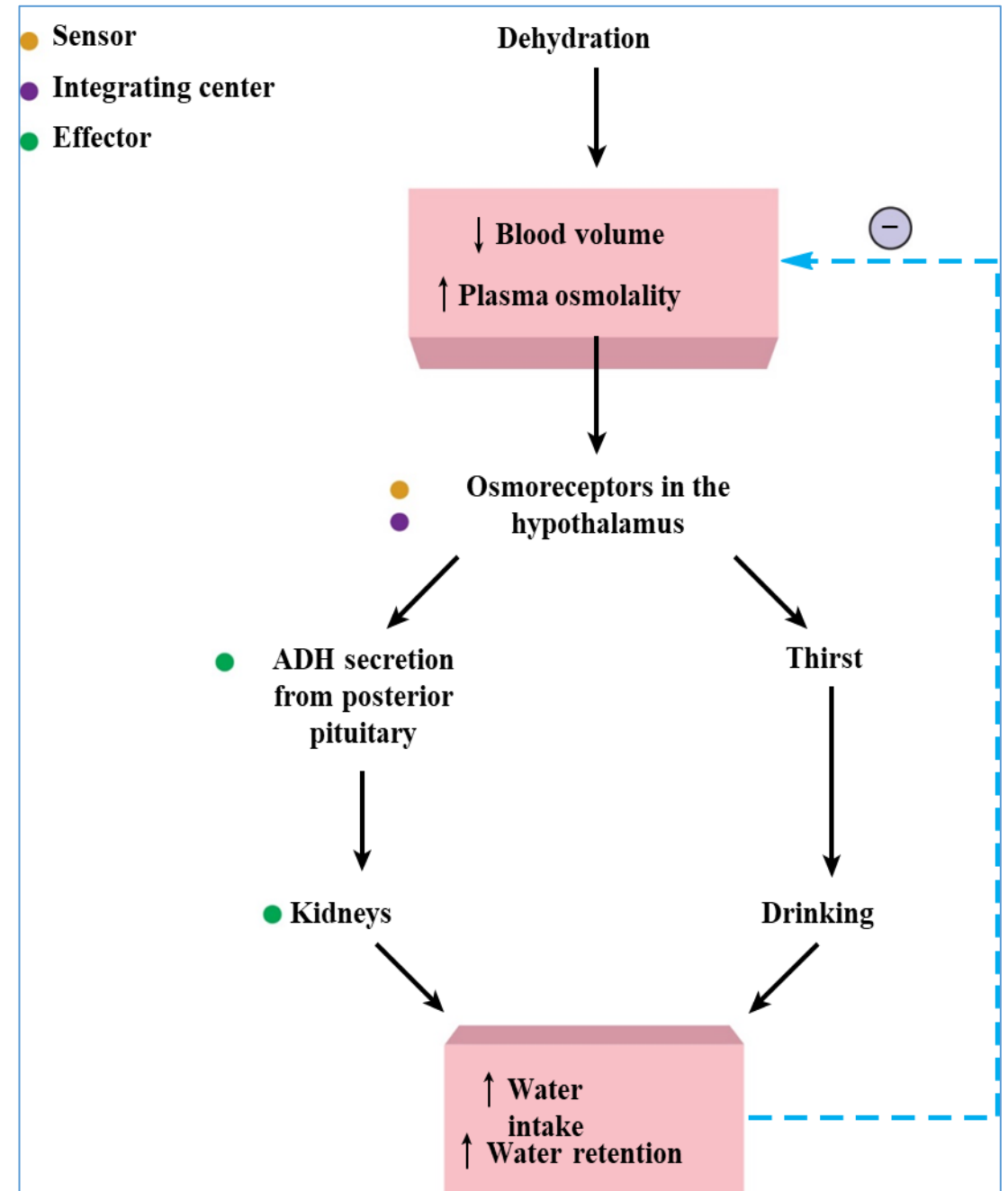
- The fate of red blood cells in isotonic, hypotonic, and hypertonic solutions.
- a) Solutions with a **lower** solute concentration than the cell are **hypoosmotic and hypotonic**.
  - a) Will pull water into the cell; cell will swell and could lyse
- b) Solutions with a **higher** solute concentration than the cell are **hyperosmotic and hypertonic**.
  - 1) Will pull water out of the cell; cell will shrivel up and could crenate



# Regulation of Blood Osmolality

- **Constant osmolality** must be maintained, or neurons will be damaged.
- **Osmoreceptors** in the hypothalamus detect increases in osmolality (due to dehydration). This triggers:
  - a. Thirst
  - b. Decreased excretion of water in urine
- With a lower plasma osmolality, osmoreceptors are not stimulated, so more water is excreted in urine

Your body has a unique way to control osmolality. When osmolality increases, it triggers your body to make antidiuretic hormone (ADH). This hormone tells your kidneys to keep more water inside your blood vessels and your urine becomes more concentrated. When osmolality decreases, your body doesn't make as much ADH.





# Carrier-Mediated Transport

- Molecules that are large or polar cannot diffuse across the membrane.
- Includes amino acids, glucose, and other organic molecules
- *Carrier proteins* within the plasma membrane move these molecules across.
- Characteristics of the carriers
  - a. They are specific to a given molecule.
  - b. Saturation – number of carriers is limited

# Facilitated Diffusion

➤ The common characteristics of **enzymes** and **carrier proteins** are:

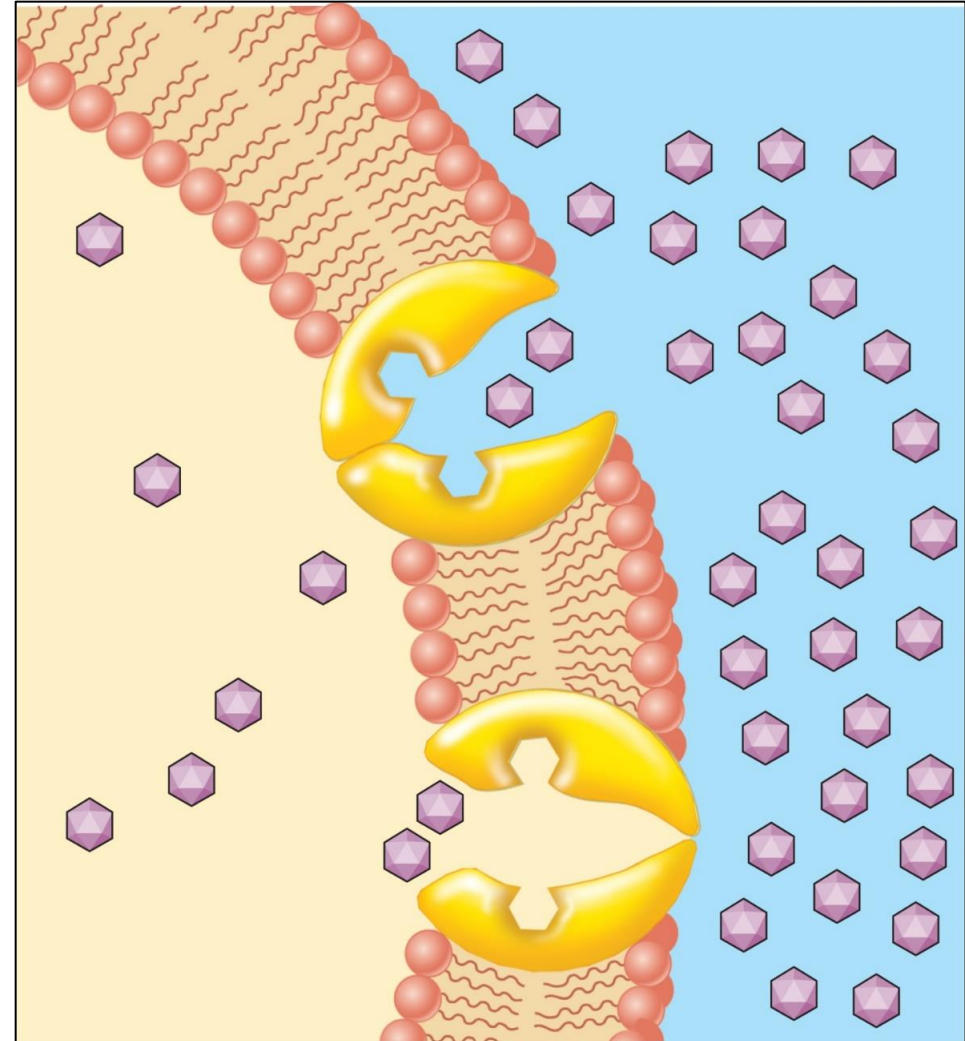
- (1) specificity
- (2) competition
- (3) Saturation

- Carrier-mediated transport that occurs against a concentration gradient, and which therefore requires metabolic energy, is called **active transport**.
- Facilitated diffusion, like simple diffusion, is powered by the thermal energy of the diffusing molecules and involves net transport from the side of higher to the side of lower concentration.
- ATP is not required for either facilitated or simple diffusion.

# Facilitated Diffusion (GLUCOSE Transport)

- The transport of glucose from the blood across plasma membranes occurs by facilitated diffusion
- ATP is not required for either facilitated or simple diffusion.
- the carrier protein has a site that can bind specifically to glucose, and such binding causes a conformational change
- The transport carriers for the facilitative diffusion of glucose are designated with the letters GLUT, followed by a number for the isoform.
- **GLUT3** in **neurons**.  
**GLUT1** is also **central nervous system**  
**GLUT2**. The **pancreatic beta** cells, **hepatocytes** of the liver

Glucose cannot move across a cell membrane via simple diffusion because it is simple large and is directly rejected by the hydrophobic tails. Instead it passes across via facilitated diffusion which involves molecules moving through the membrane by passing through channel proteins.

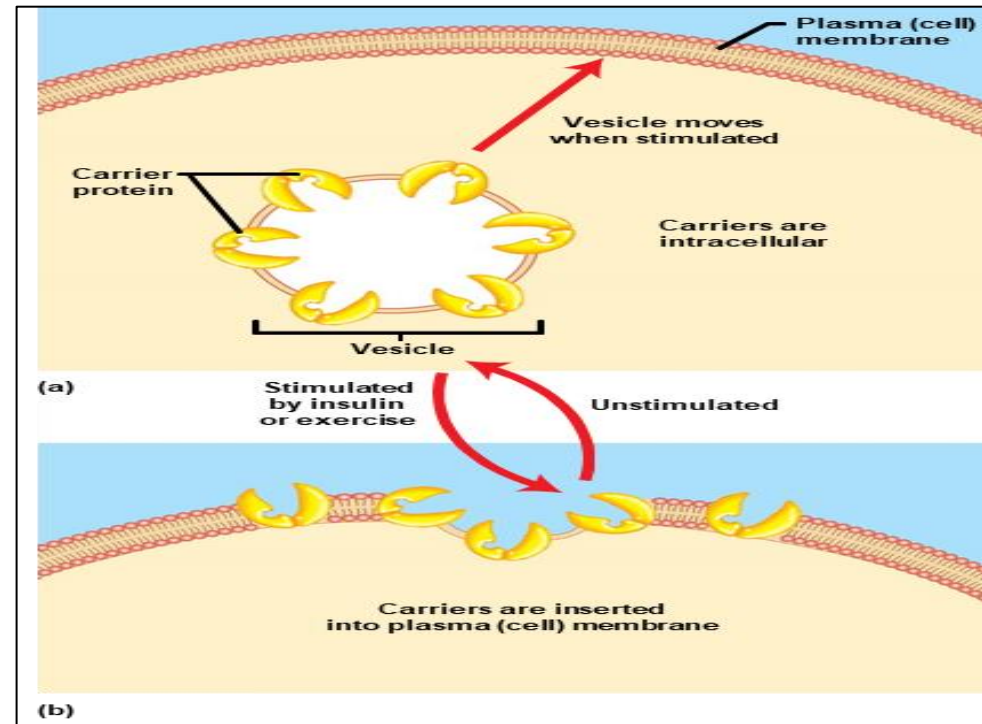


*Facilitated Diffusion of Glucose*

## Insertion of Carrier Proteins into the Plasma Membrane

Transport proteins may always exist in the plasma membrane or be inserted when needed

In unstimulated **muscles**, the **GLUT4** proteins are located within the membrane of cytoplasmic vesicles. Exercise—and stimulation by insulin—causes these vesicles to fuse with the plasma membrane.



# Diffusion through the Plasma Membrane

## Non carrier-mediated

**Simple diffusion**  
of lipid soluble  
molecules through  
the phospholipid  
bilayer  
ex. O<sub>2</sub>, CO<sub>2</sub>  
steroid hormones

**Simple diffusion**  
of ions through  
membrane channel  
proteins  
ex. Na<sup>+</sup>, Cl<sup>-</sup>

**Simple diffusion**  
of water molecules  
(osmosis) through  
aquaporin (water)  
channels

## Carrier-mediated

**Facilitated  
diffusion**  
needs carrier to  
pass through the  
plasma membrane

**Active transport**  
needs ATP  
against the  
concentration  
gradient  
through Pumps

passive transport

# Active Transport

- Sometimes molecules must be moved from an area of low concentration to an area of high concentration (move uphill):
  - a. This requires the expenditure of ATP.
  - b. Often, these carrier-mediated proteins are called pumps.



# Active Transport

```
graph TD; A[Active Transport] --> B[Primary Active Transport]; A --> C[Secondary Active Transport]; B --> D[The hydrolysis of ATP is directly responsible for the function of the carriers]; D --> E[ex. Ca2+ Pump, H+ pump, Na / K ATPase]; C --> F[The energy needed for the “uphill” movement of a molecule or ion is obtained from the “downhill” transport of Na+ into the cell]; F --> G[ex. Cotransport/symport: Glucose and Na+ in kidney and intestine lumen<br/>ex. Countertransport/Antiport: Na+ / Ca 2+];
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## Primary Active Transport

The hydrolysis of ATP is directly responsible for the function of the carriers

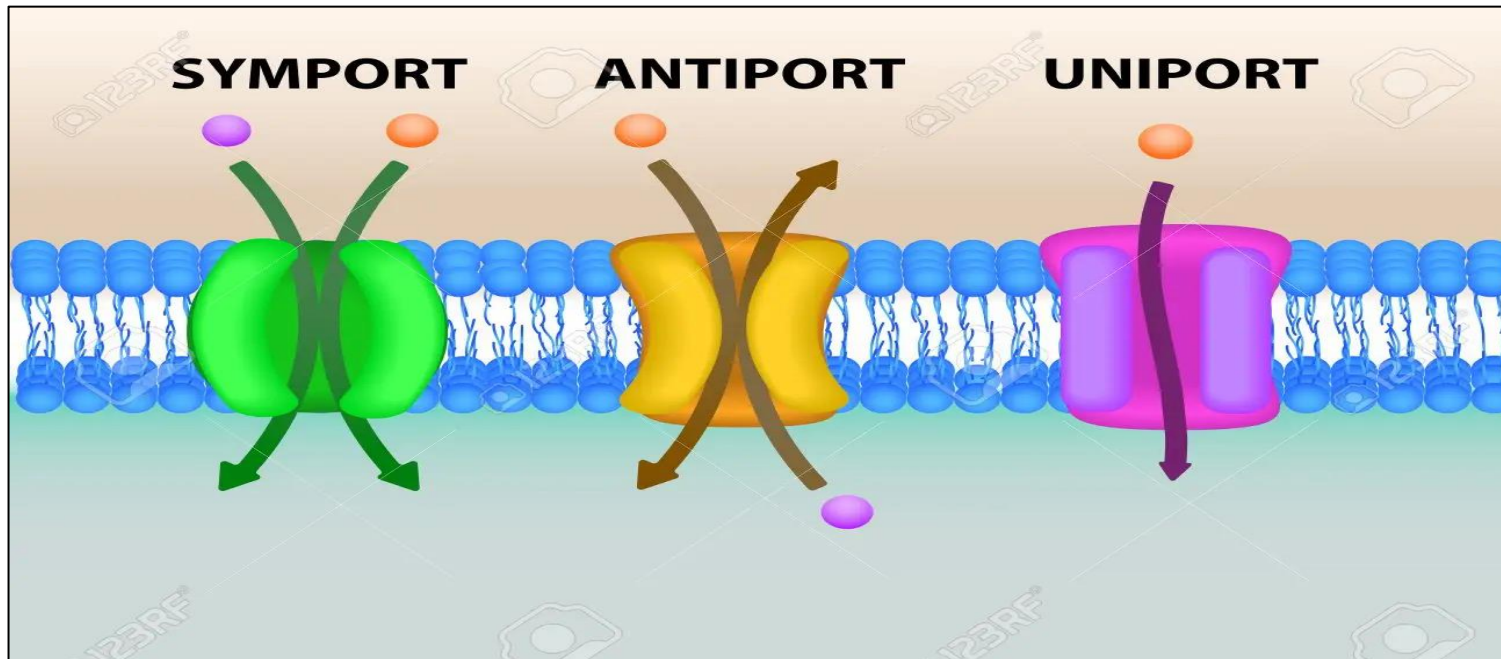
ex.  $\text{Ca}^{2+}$  Pump,  $\text{H}^+$  pump, Na / K ATPase

## Secondary Active Transport

The energy needed for the “uphill” movement of a molecule or ion is obtained from the “downhill” transport of  $\text{Na}^+$  into the cell

ex. Cotransport/symport: Glucose and  $\text{Na}^+$  in kidney and intestine lumen  
ex. Countertransport/Antiport:  $\text{Na}^+$  /  $\text{Ca}^{2+}$

- **A uniport** is the transport of only one molecule, without coupling to the transport of another molecule or ion.
- **Symport** is the type of transport in which two different molecules can move through a membrane in same direction by using a common carrier mechanism.
- **Antiport** is a type of transport in which two different molecules can move through a membrane in opposite directions.

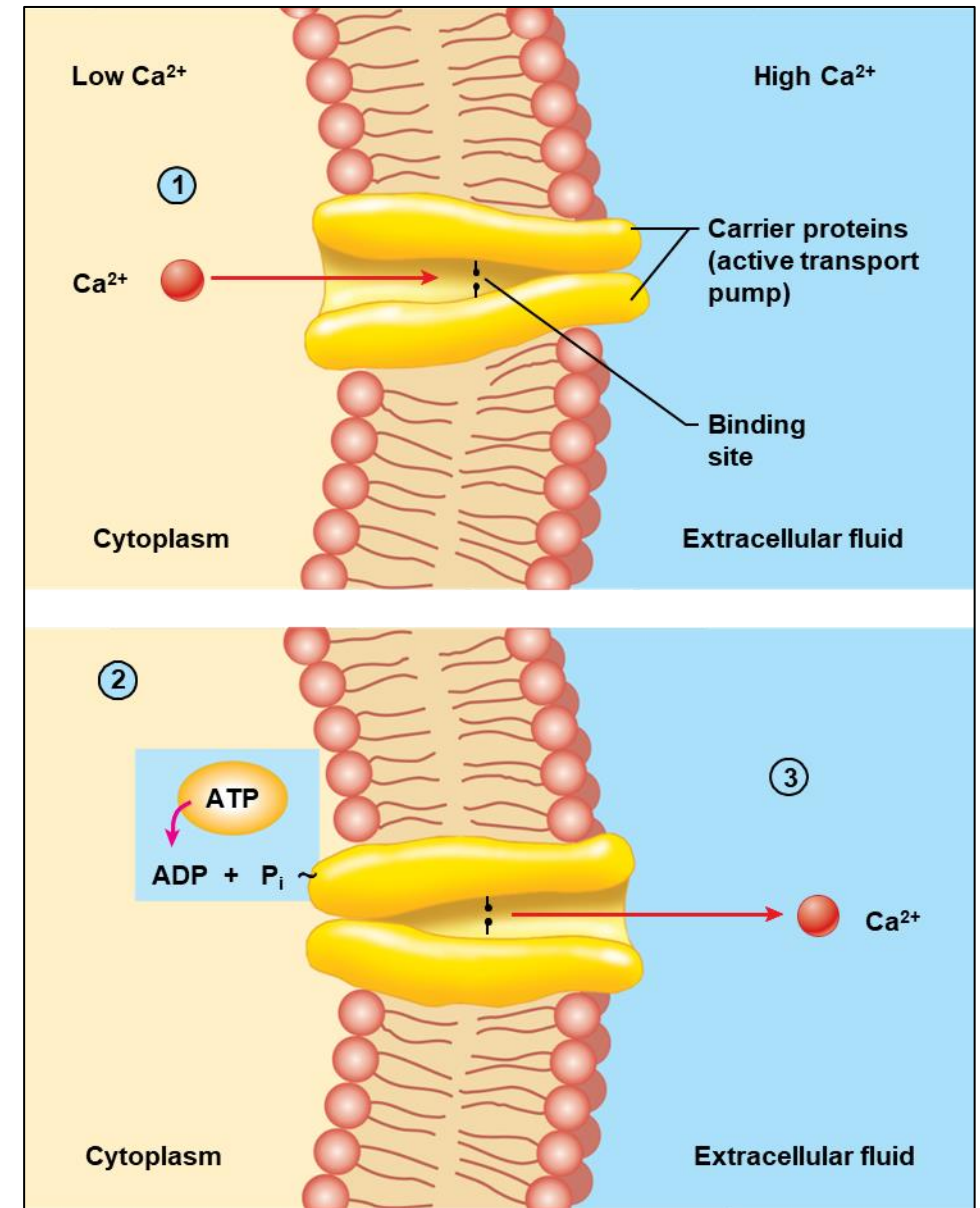


# Primary Active Transport

- a. Occurs when the hydrolysis of ATP is directly responsible for the carrier protein function.
- b. The transport protein is also an **ATPase** enzyme that will hydrolyze ATP.
- c. Pump is activated by **phosphorylation** using a  $P_i$  from ATP.

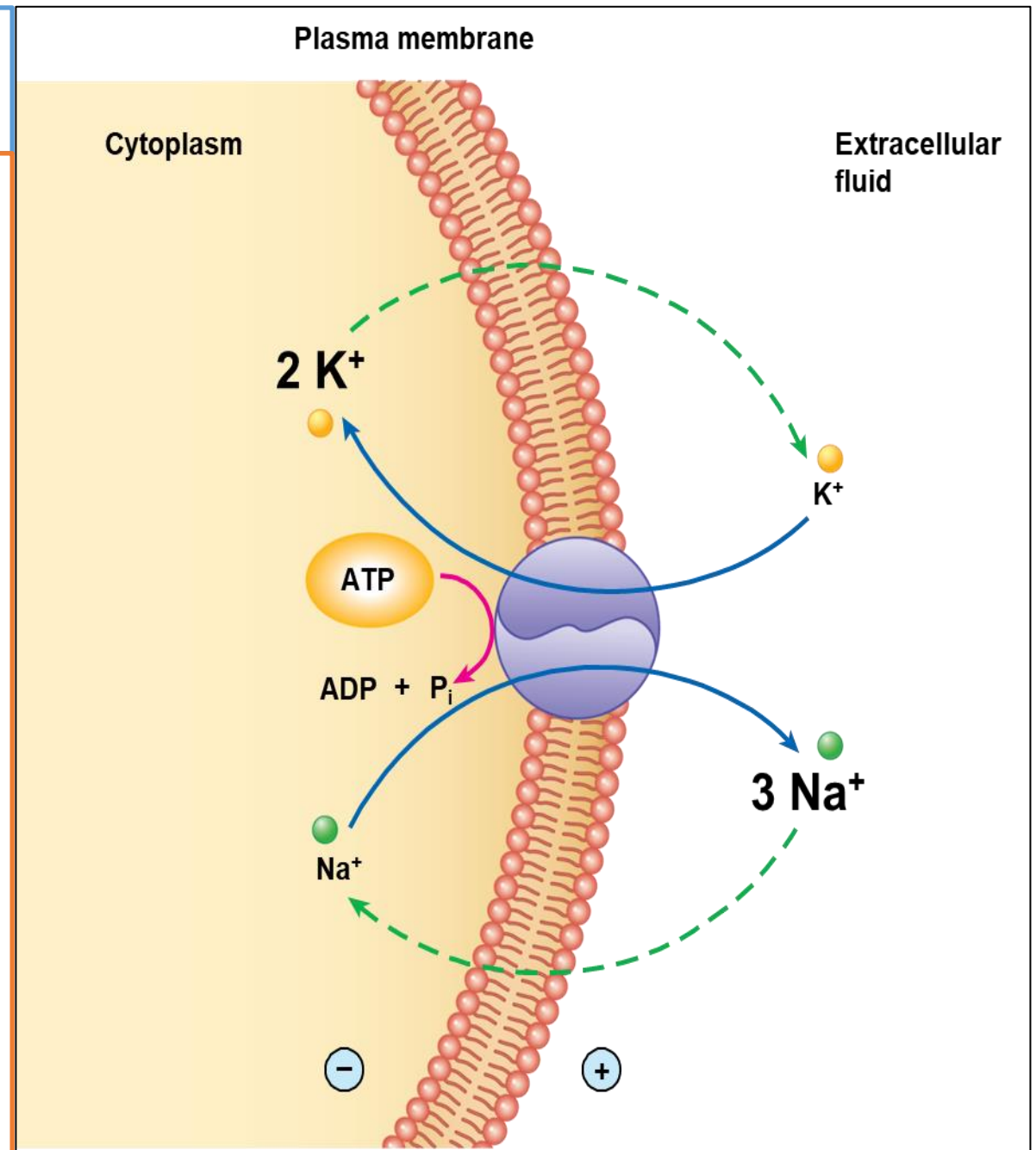
# The $\text{Ca}^{2+}$ Pump

- a. Located on all cells PM and in the endoplasmic reticulum (ER) of striated muscle cells
- b. Removes  $\text{Ca}^{2+}$  from the **cytoplasm** by pumping it into the extracellular fluid or **cisternae of the ER**
- c. Creates a strong **concentration gradient** for rapid movement of  $\text{Ca}^{2+}$  back into the cell (**signal**)
- d. Aids in release of **neurotransmitters** in neurons and in **muscle contraction**



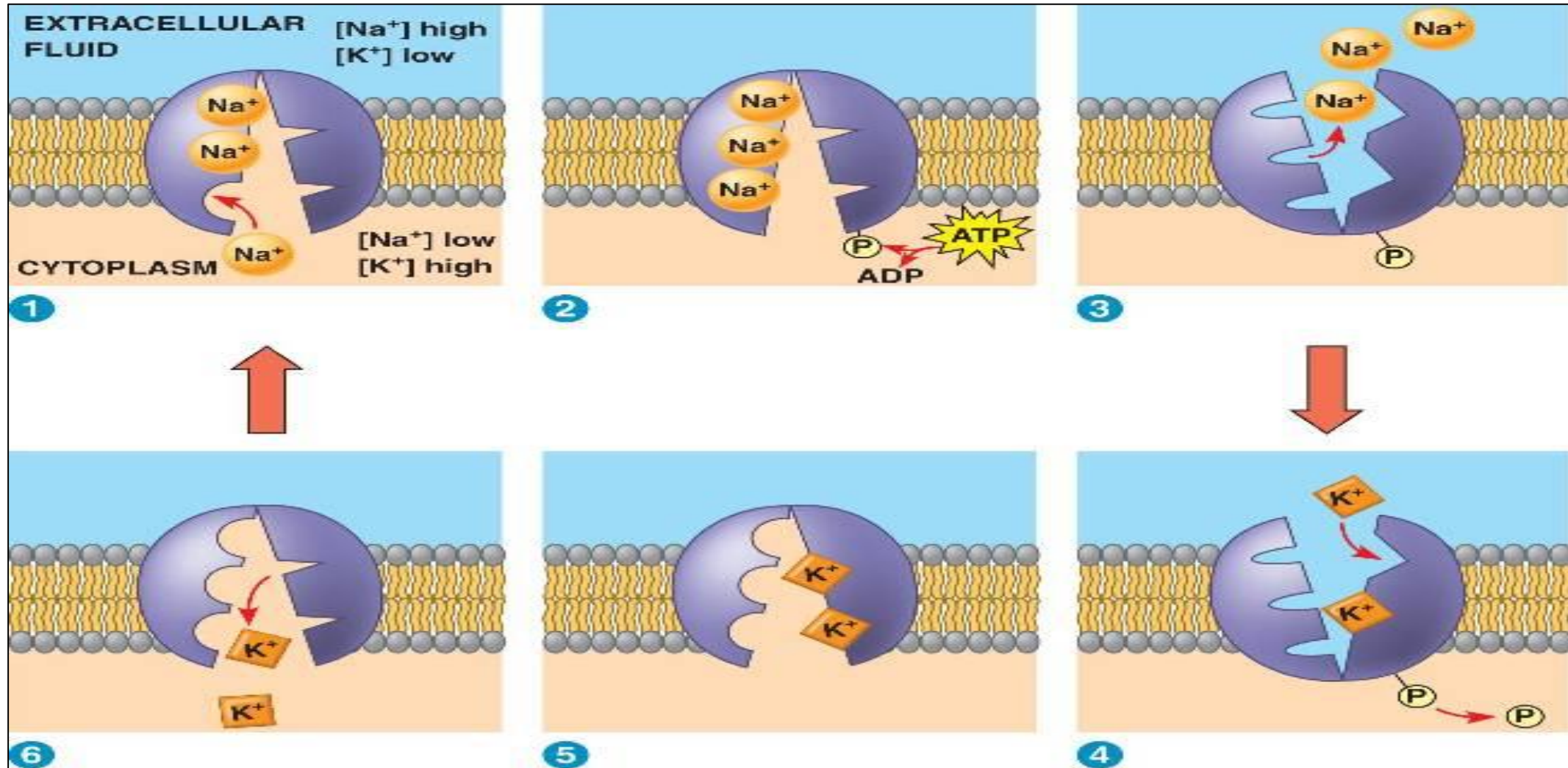
# Na<sup>+</sup>/K<sup>+</sup> Pump

- a. Found in all body cells
  - b. ATPase enzyme pumps 3 Na<sup>+</sup> out of the cell and 2 K<sup>+</sup> into the cell
- The steep gradient of Na<sup>+</sup> and K<sup>+</sup> across the plasma membrane serves three functions:
    - 1) Provides **energy** for coupled transport of other molecules.
    - 2) Produces **electrochemical impulses** in neuron and muscle cells including the heart muscle.
    - 3) Maintains **osmolality** (otherwise increased Na<sup>+</sup> concentrations within cells would promote the osmotic inflow of water, damaging the cells).





# Na<sup>+</sup>/K<sup>+</sup> Pump





## Secondary Active Transport of Glucose

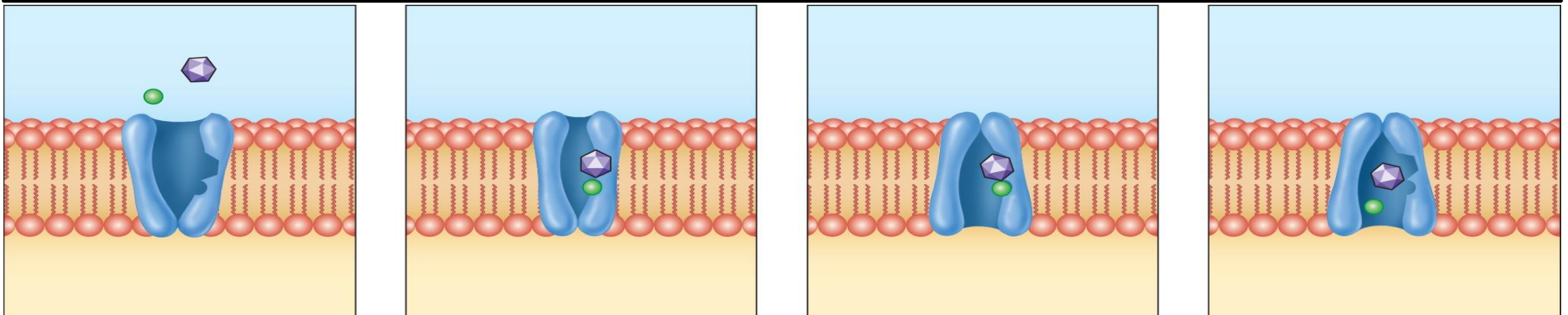
### The Co-transport of $\text{Na}^+$ and glucose.

This carrier protein transports Na and glucose at the same time, moving them from the lumen of the intestine and kidney tubules into the lining epithelial cells.

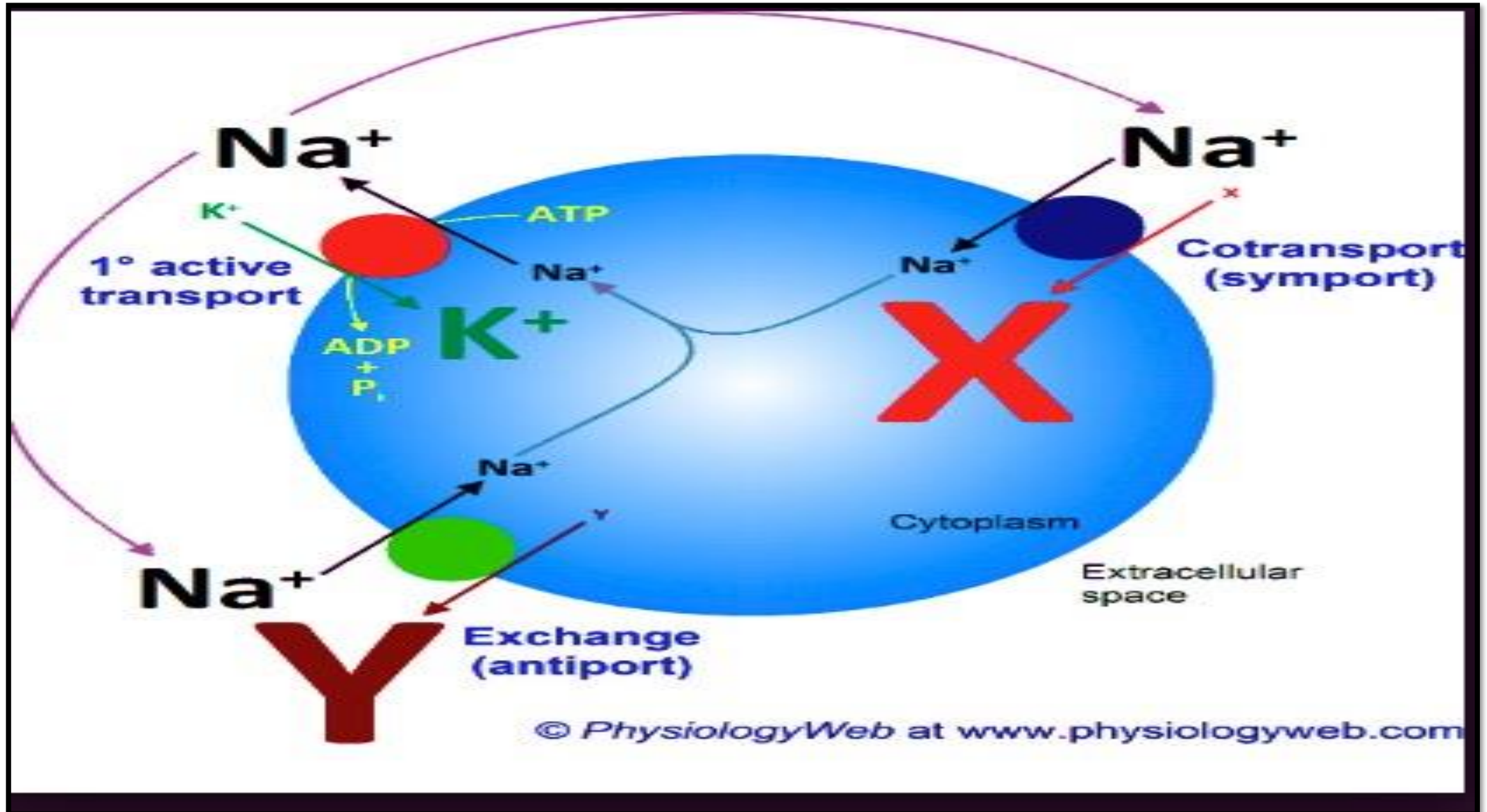
This co-transport requires a lower intracellular concentration of  $\text{Na}^+$ , which is dependent on the action of other carriers, the  $\text{Na}^+/\text{K}^+$  (ATPase) pumps.

Because ATP is needed to power the  $\text{Na}^+/\text{K}^+$  (ATPase) pumps, the co-transport of Na and glucose depends indirectly on ATP, and so can be considered secondary active transport.

The co-transport carrier shown here transports 1 Na to 1 glucose, as most commonly occurs in the kidney; the carrier in the small intestine transports 2 Na 1 for 1 glucose (not shown).



# Secondary Active Transport

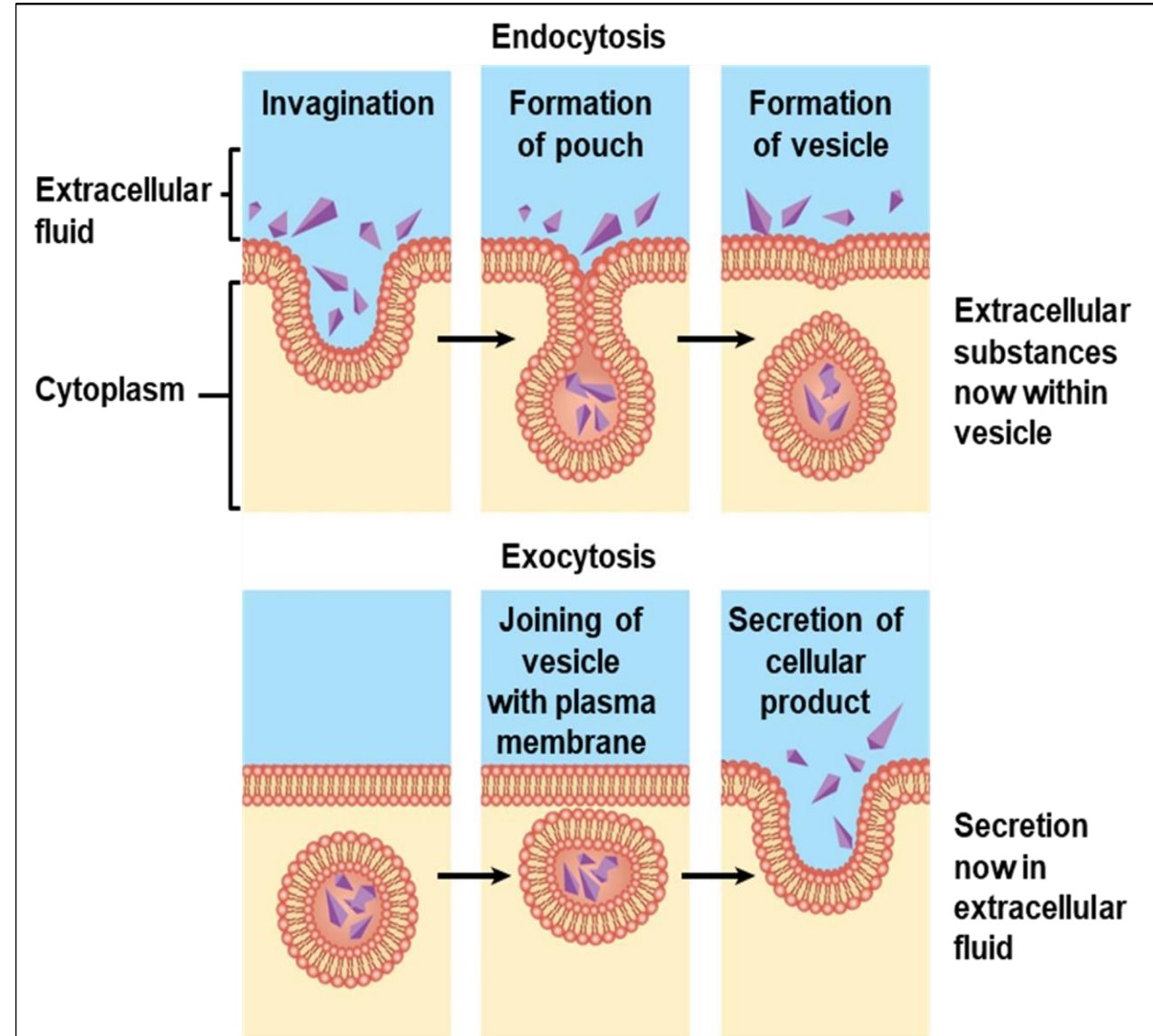


# Bulk Transport

- Large molecules such as **proteins, hormones, and neurotransmitters** are secreted via **exocytosis**.
  - Involves fusion of a vesicle with the plasma membrane
  - Requires ATP
- Movement of large molecules such as cholesterol into the cell requires ***endocytosis***.
  - Usually a transport protein interacts with plasma membrane proteins to trigger endocytosis.

# Bulk Transport

- Large molecules such as **proteins, hormones, and neurotransmitters** are secreted via **exocytosis**.
- Involves fusion of a vesicle with the plasma membrane
- Requires ATP
- Movement of large molecules such as cholesterol into the cell requires **endocytosis**.
- Usually a transport protein interacts with plasma membrane proteins to trigger endocytosis.



# The Membrane Potential

1. There is unequal distribution of charge across the plasma membrane due to
  - a. Permeability properties of the plasma membrane
  - b. Action of  $\text{Na}^+/\text{K}^+$  pumps
  - c. Negatively charged molecules inside the cell
2. This difference in charge is called a **potential difference**
3. The potential difference makes the inside of the cell negative compared to the outside (**Cellular proteins, phosphate groups of ATP, other organic molecules are negatively charged at the pH of the cell cytoplasm**)



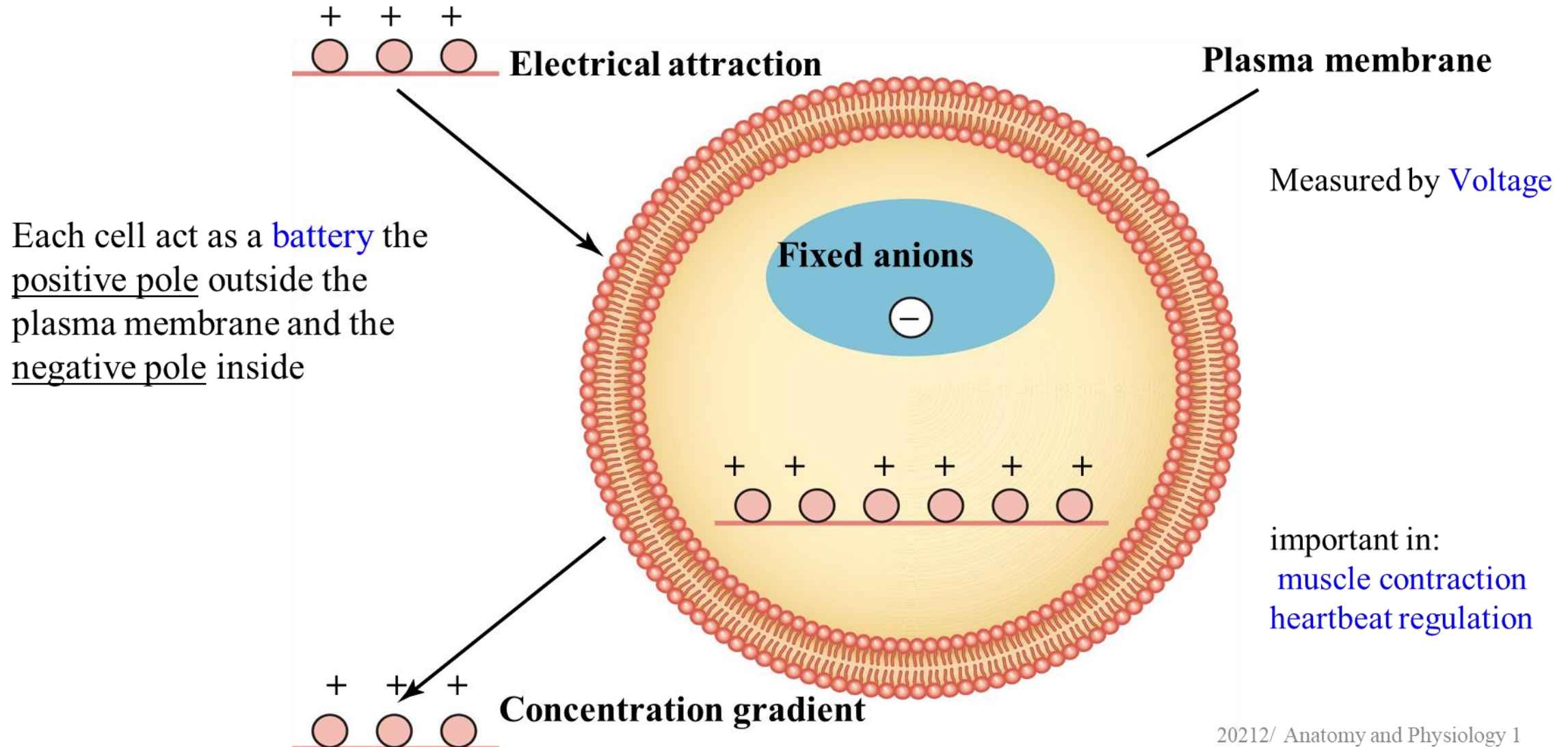
# The Membrane Potential

- These **negative ions ( anions )** are “**fixed**” within the cell because they cannot penetrate the plasma membrane.
- As a result, these anions attract positively charged inorganic ions (**cations**) from the extracellular fluid that can pass through ion channels in the plasma membrane
- Fixed anions within the cell influence the distribution of inorganic cations (mainly  $K^+$ ,  $Na^+$ , and  $Ca^{++}$ ) between the extracellular and intracellular compartments.



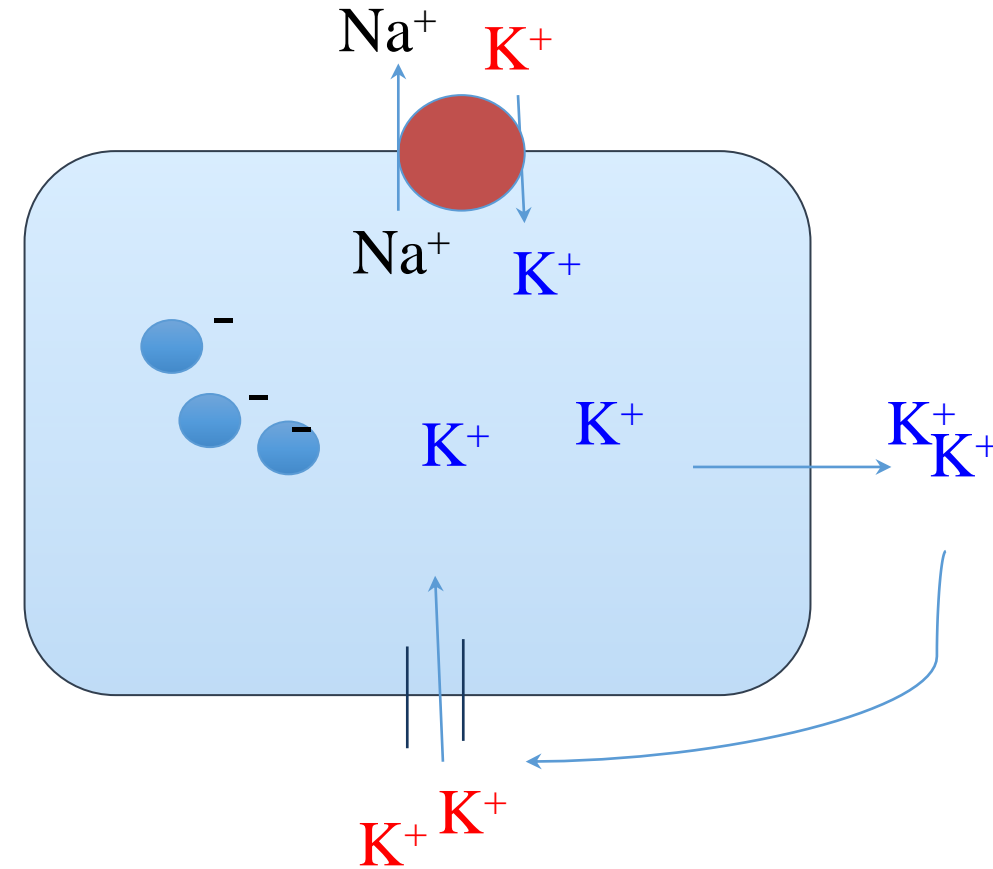
# Effect of Fixed Anions on Distribution of Cations

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# Membrane Potential: $K^+$

- $K^+$  accumulates at high concentrations in the cell because:
  - a. The  $Na^+/K^+$  pumps actively bring in  $K^+$
  - b. The membrane is very permeable to  $K^+$
  - c. Negative anions inside the cell attract cations outside the cell.
  - d. Limited by strong concentration gradient.
- The  $K^+$  concentration inside is 150 mEq/L and out is 5 mEq/L

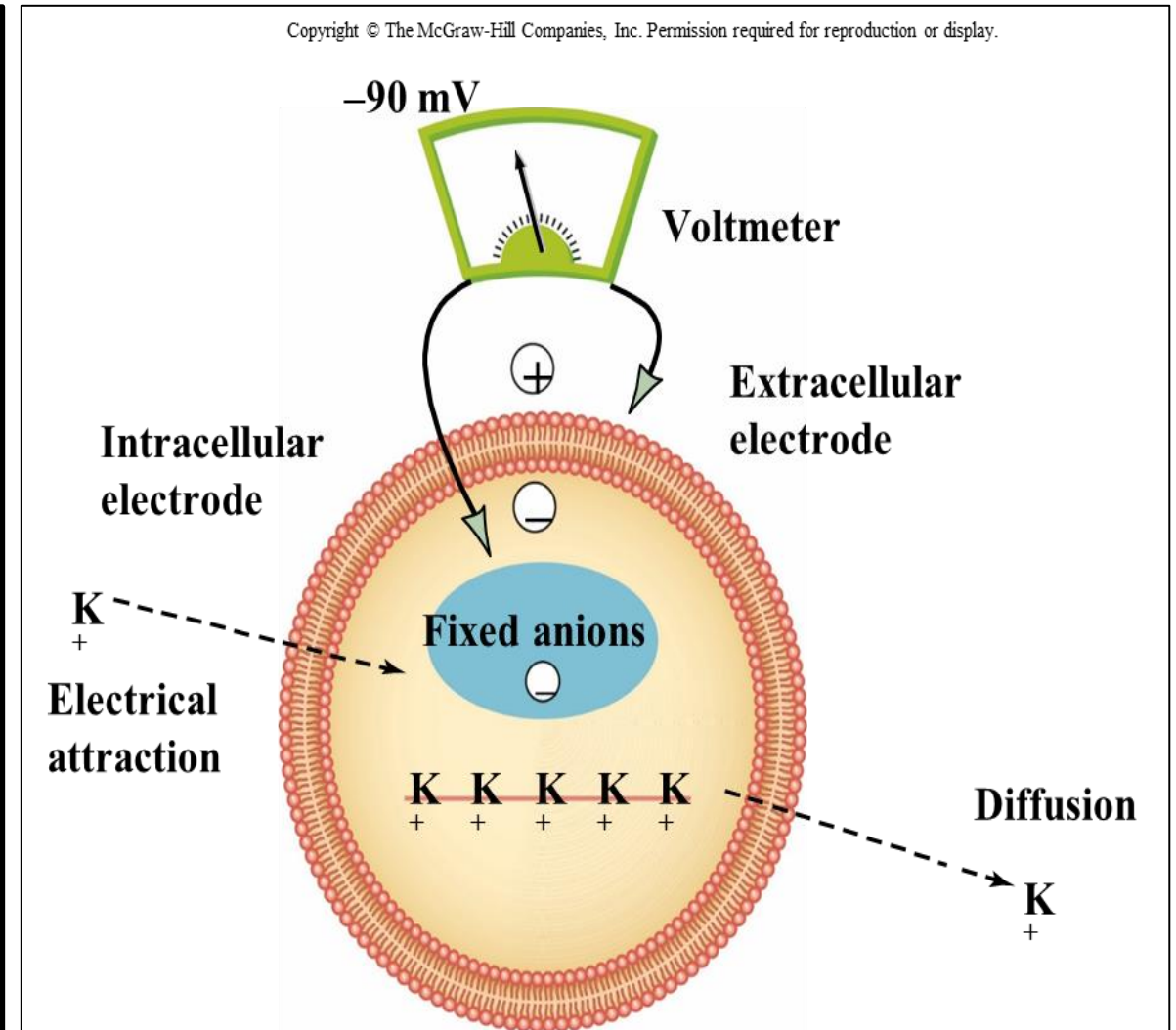


# Equilibrium Potentials

- The Plasma membrane is **more permeable** to  $K^+$  than other ions, the membrane potential is determined by  $K^+$  concentration gradient.
- Even with all the  $K^+$  inside the cell, the negative molecules inside and all of the sodium outside, **the cell is more negative inside compared to outside.**

# K<sup>+</sup> Equilibrium Potential

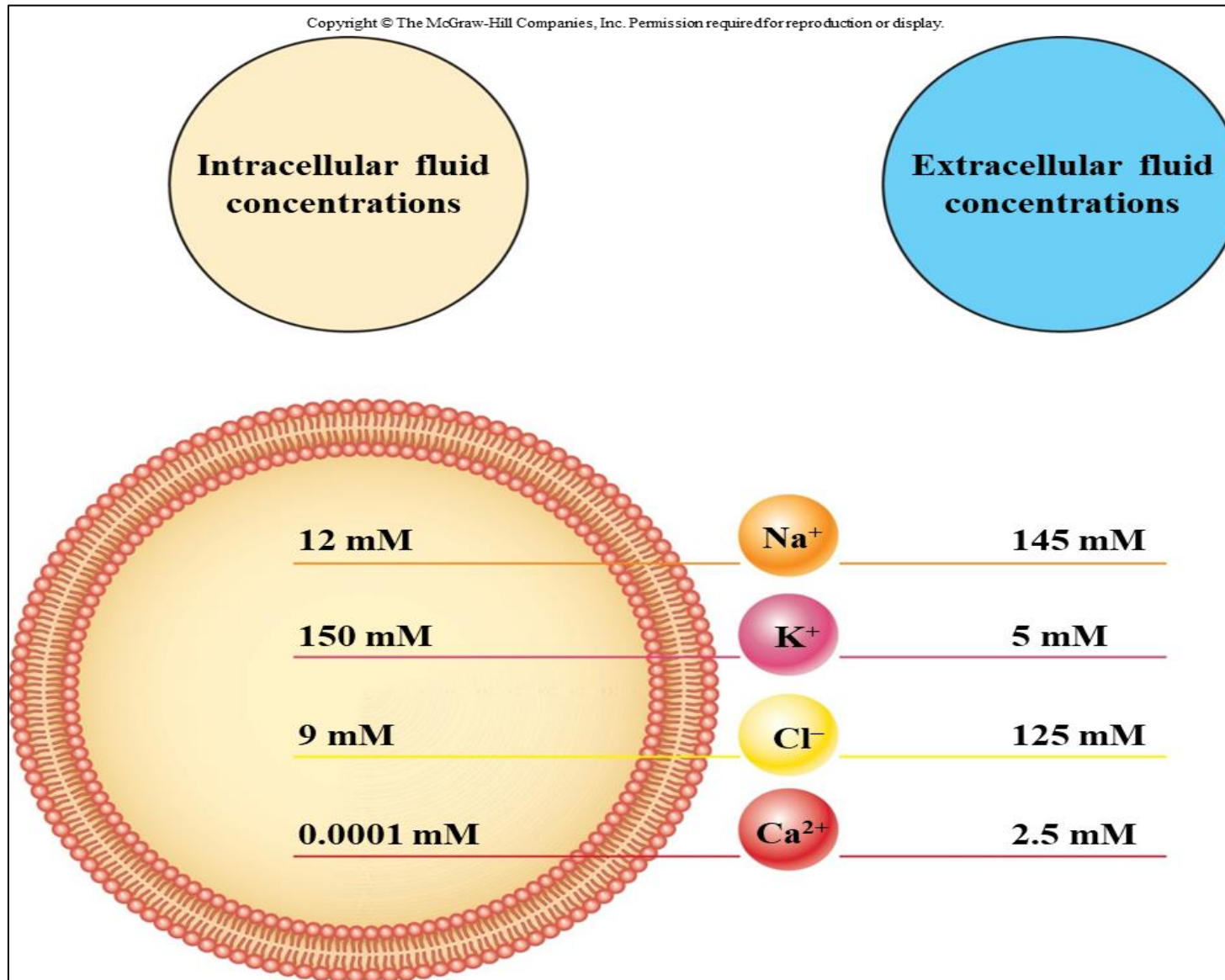
- **Addressing just K<sup>+</sup>**, the electrical attraction would pull K<sup>+</sup> into the cell until it reaches a point where the concentration gradient drawing K<sup>+</sup> out matches this pull in.
  - 1) K<sup>+</sup> would reach an equilibrium, with more K<sup>+</sup> inside than outside.
  - 2) Normal cells have 150 mM K<sup>+</sup> inside and 5mM K<sup>+</sup> outside
- **The resulting potential difference** measured in voltage would be the equilibrium potential (E<sub>K</sub>) for K<sup>+</sup>; measured at -90mV.
  - 1) This means the inside has a voltage 90mV lower than the outside.
  - 2) This is the voltage needed to maintain 150 mM K<sup>+</sup> inside and 5mM K<sup>+</sup> outside.
  - ❖ A state of equilibrium would be reached where the concentrations of K<sup>+</sup> remained stable



# Na<sup>+</sup> Equilibrium

- A. Sodium is also an important ion for establishing membrane potential.
- B. The concentration of sodium in a normal cell is 12mM inside and 145mM outside.
- C. To keep so much sodium out, the inside would have to be positive to repel the sodium ions.
- D. The equilibrium potential for sodium is +66mV.
- E. The membrane is less permeable to Na<sup>+</sup>, so the actual membrane potential is closer to that of the more permeable K<sup>+</sup>.

# Concentration of ions in the intracellular and extracellular fluids





# Resting Membrane Potential

- **Resting Membrane Potential(RMP)**: Membrane potential of a cell not producing any impulses.
- Depends on:
  - a. Ratio of the concentrations of each ion on either side of the membrane
  - b. Specific permeability to each ion ( $K^+$ ,  $Na^+$ ,  $Ca^{2+}$  and  $Cl^-$  contribute to the resting potential)
- If the plasma membrane were only permeable to  $Na^+$  , its resting membrane potential would equal the  $E_{Na}$  of +66 mV; if it were only permeable to  $K^+$ , its resting membrane potential would equal the  $E_K$  of -90 mV.

# Resting Membrane Potential

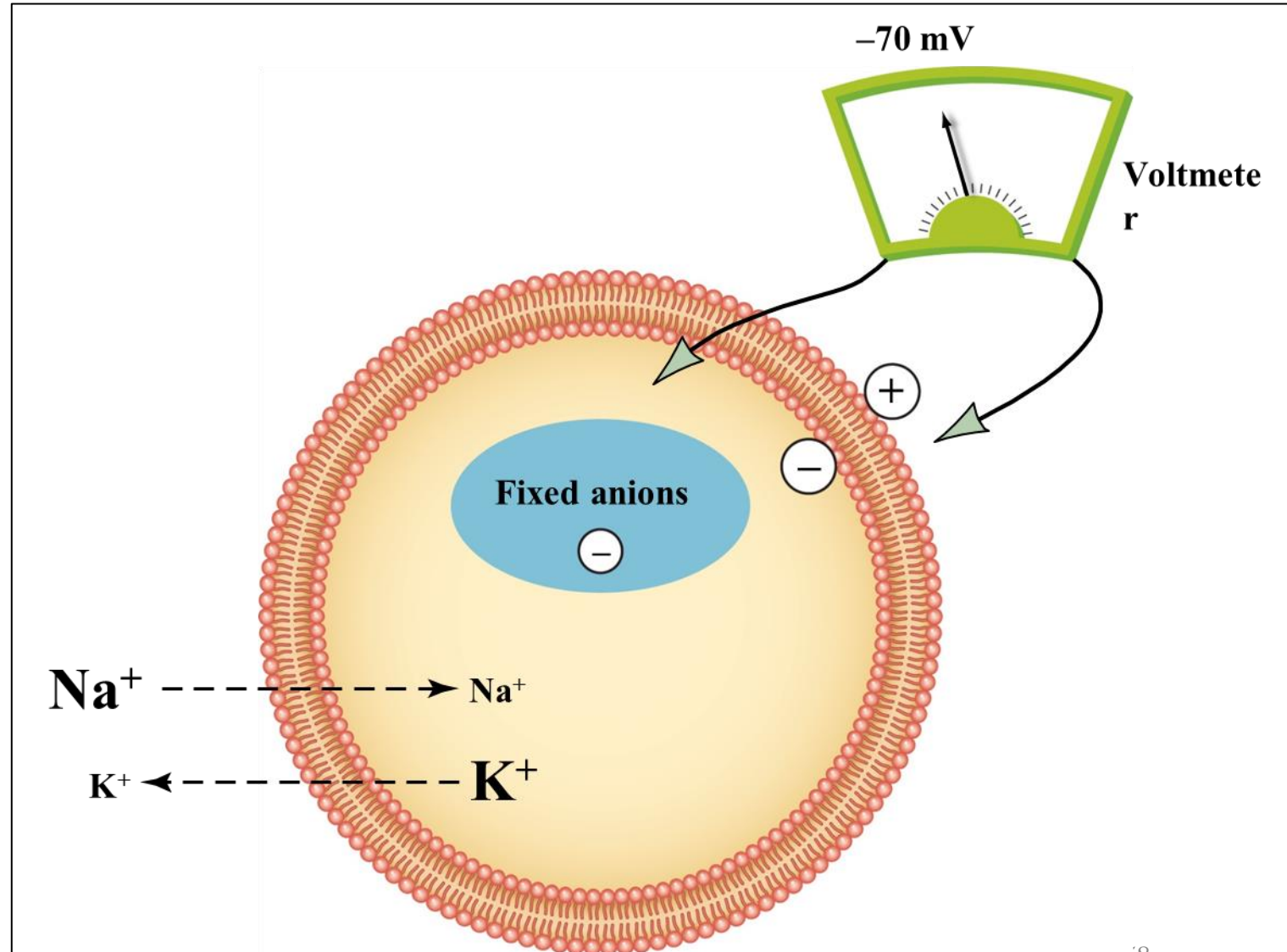
- The concentration of ions contributes to the (RMP):
  - a. Because the membrane is most permeable to  $K^+$ , it has the greatest influence.
  - b. A change in the permeability of the membrane for any ion will change the resting potential.
  - c. A change in the concentration of any ion inside or outside the cell will change the resting potential

Key to how neurons work

- In most cells, the resting potential is between -65mV and -85mV.
  - a. Neurons resting membrane potential is usually  $-70\text{mV}$ .
  - b. Close to  $K^+$  equilibrium potential
- When a **neuron sends an impulse**, it changes the permeability of  $Na^+$ , driving the membrane potential closer to the equilibrium potential for  $Na^+$

# Resting Membrane Potential

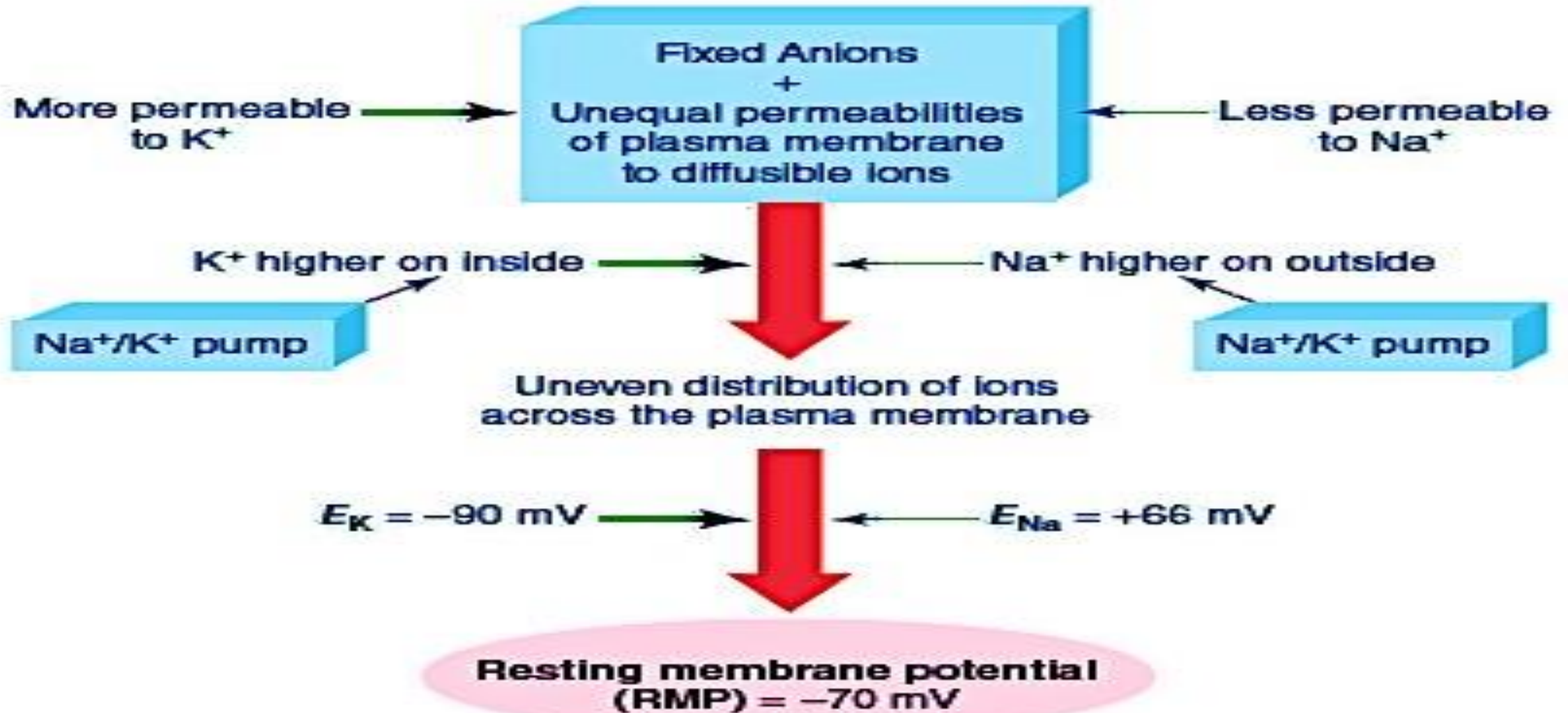
Because some  $\text{Na}^+$  leaks into the cell by diffusion, the actual resting membrane potential is not as negative as the  $\text{K}^+$  equilibrium potential. As a result, some  $\text{K}^+$  diffuses out of the cell, as indicated by the dashed lines.



# Role of Na<sup>+</sup>/K<sup>+</sup> Pump

- Acts to counter K<sup>+</sup> and Na<sup>+</sup> leaking out at the resting potential.
- It transports 2 K<sup>+</sup> in for every 3 Na<sup>+</sup> out to maintain the voltage difference.
- Contribute to the negative intracellular charge (Electrogenic effect) Keeps both the resting potential and the concentration differences stable

# Processes that influence the resting membrane potential

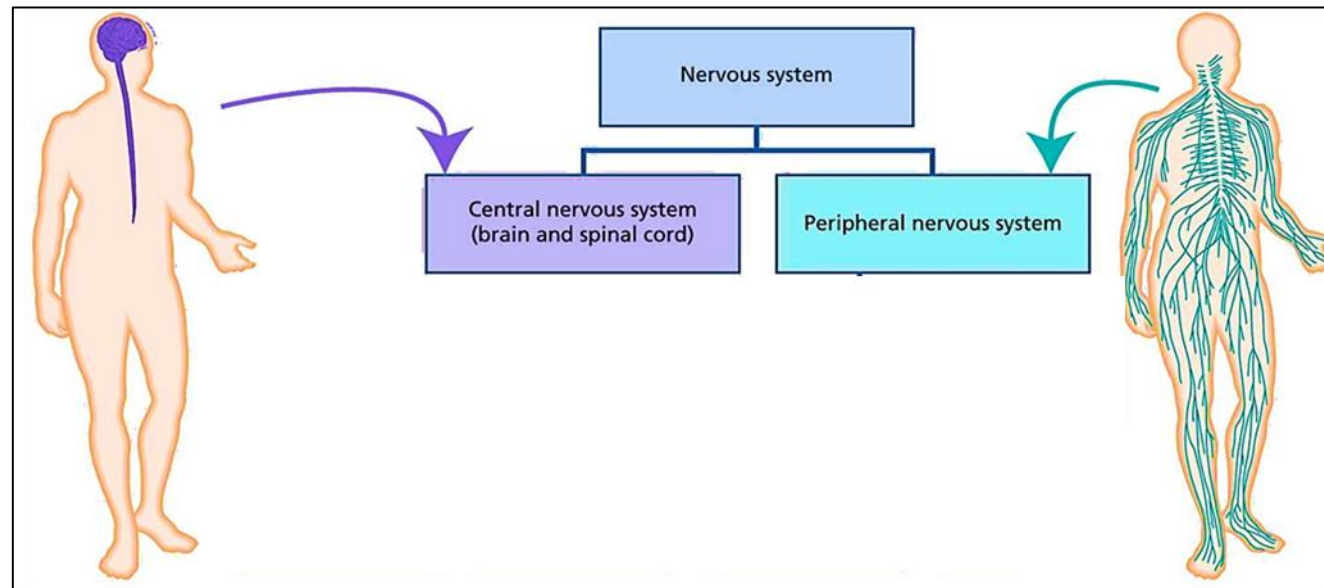


# **Chapter 7: Introduction to the Nervous System**



# Chapter 7: Introduction to the Nervous System

- Divided into:
  - a. **Central nervous system:** brain and spinal cord
  - b. **Peripheral nervous system:** cranial nerves (arising from the brain), and spinal nerves (arising from the spinal cord)
- Nervous tissue is composed of two types of cells:
  - a. **Neurons** that conduct impulses but generally can not divide.
  - b. **Glial cells (neuroglia)** that support the neurons and can not conduct impulses but can divide.



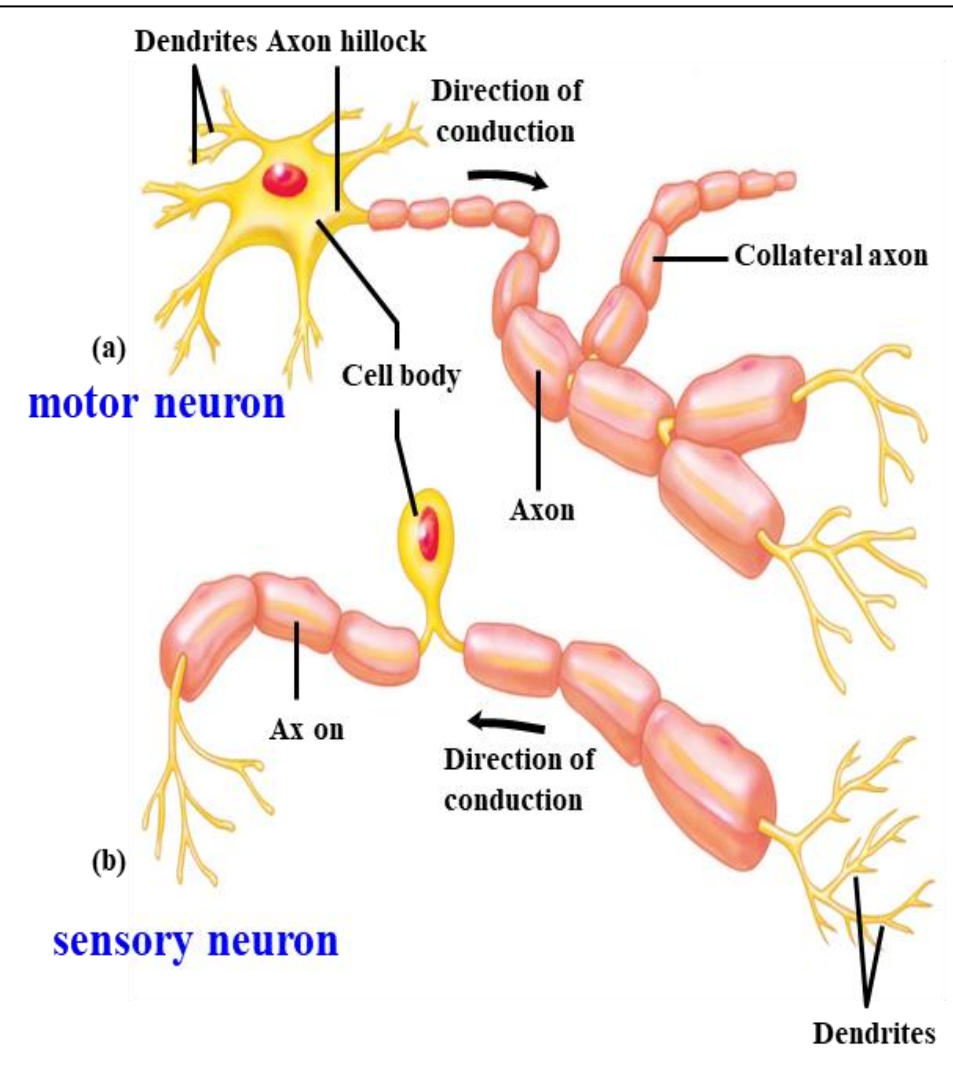
# Neurons

- The basic structural and functional units of the nervous system.
- General functions
  - a. Respond to chemical and physical stimuli
  - b. Conduct electrochemical impulses
  - c. Release chemical regulators
  - d. Enable perception of sensory stimuli, learning, memory, and control of muscles and glands
- Most can not divide, but can repair.
  - Unlike neurons glial cells which are able to divide by mitosis.
  - This helps to explain why brain tumors in adults are usually composed of glial cells rather than of neurons.

# General structure of neurons

Neurons vary in size and shape, but they all have:

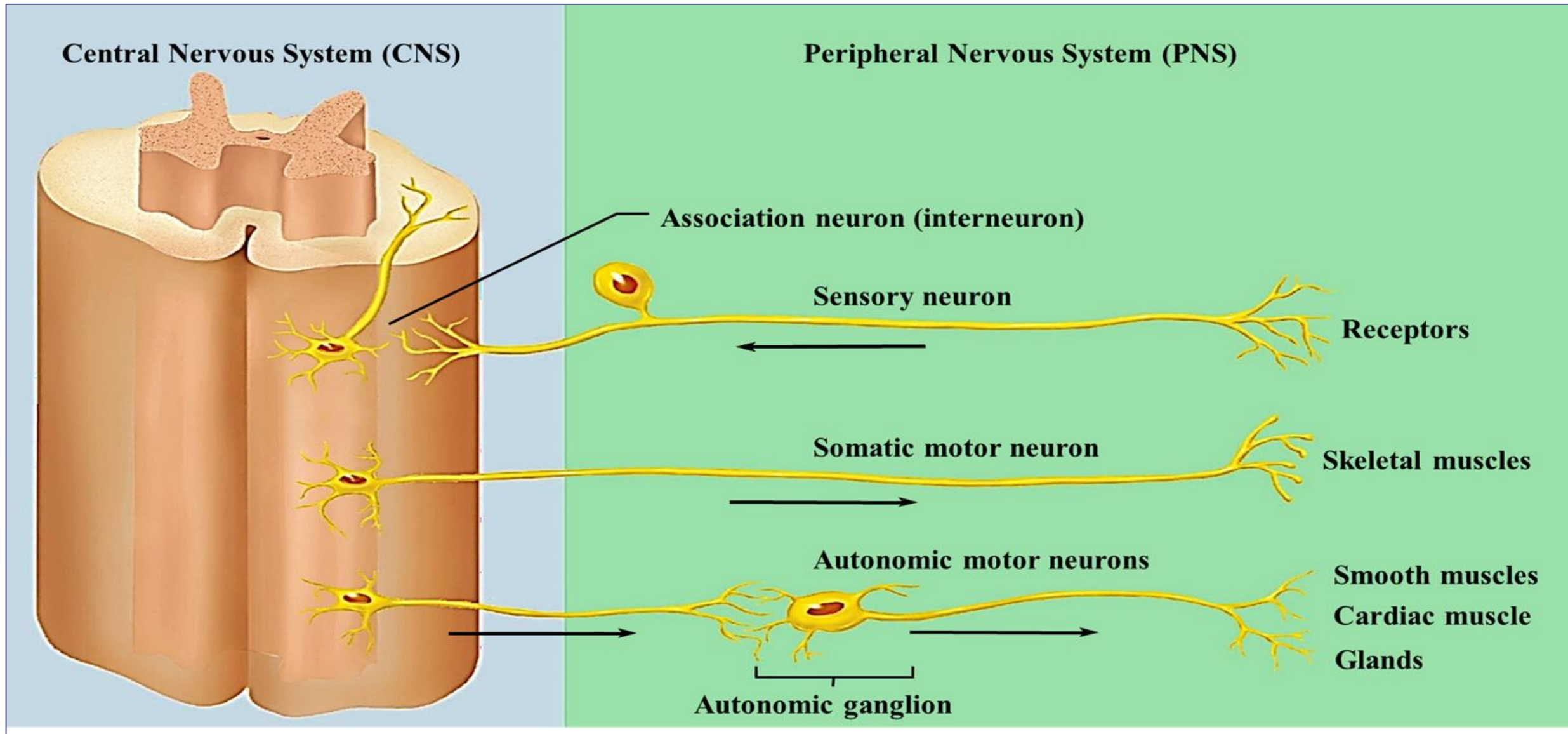
1. **A cell body:** that contains the nucleus, Nissl bodies (large stacks of rough endoplasmic reticulum that are needed for the synthesis of membrane proteins); they cluster in groups called *nuclei* in the CNS and *ganglia* in the PNS.
2. **Dendrites:** receive impulses and conducts a graded impulse (local event) toward the cell body.
3. **Axon:** conducts action potentials away from the cell body.
  - Vary in length from a few millimeters to a meter
  - Connected to the cell body by the axon hillock (the origin of the axon) where action potentials are generated at the initial segment of the axon.
  - Covered in myelin with open spots called nodes of Ranvier



# Functional classification of Neurons

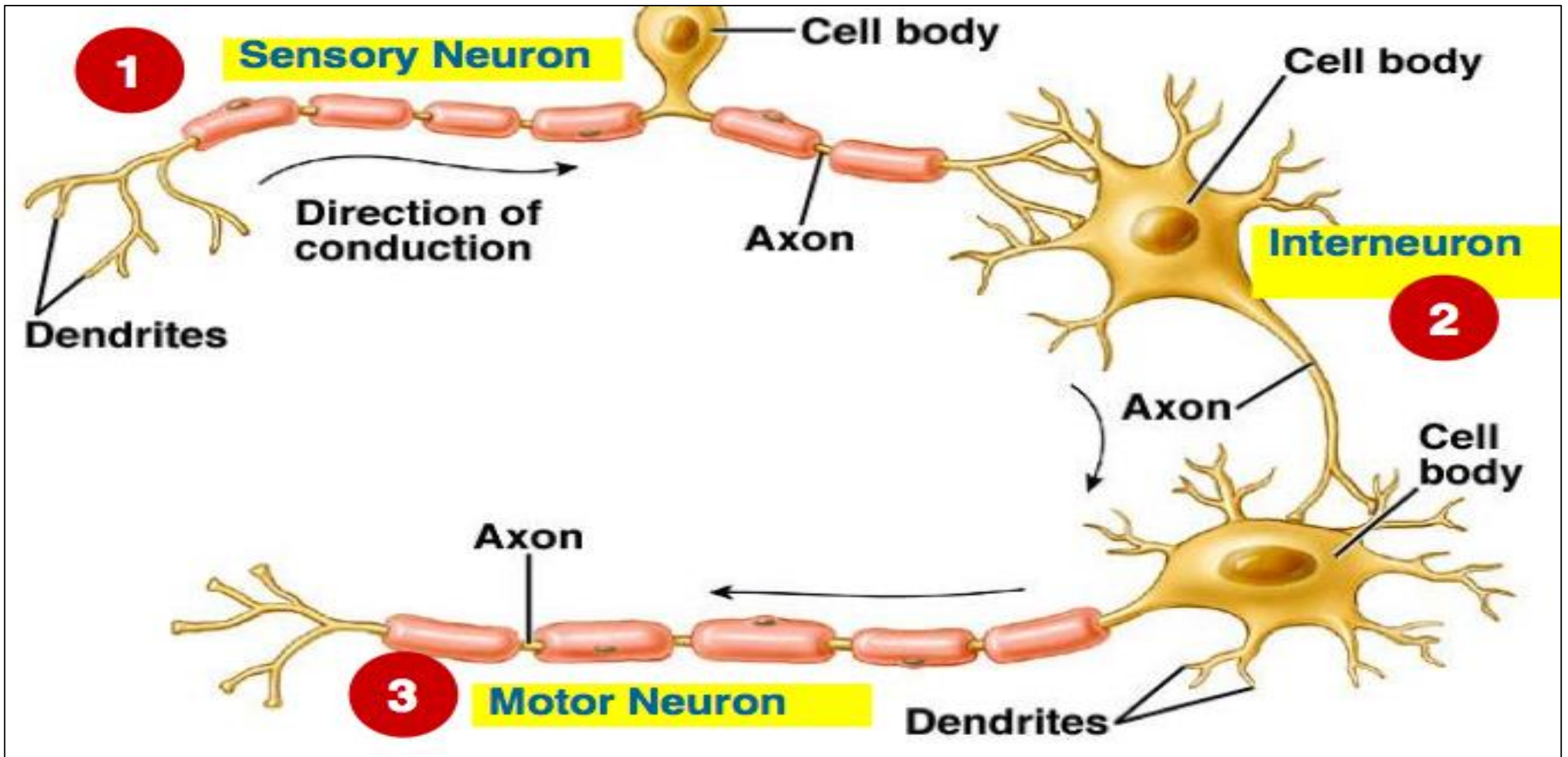
1. Functional classification of neurons – based on direction impulses are conducted.
  1. **Sensory or afferent neurons** : conduct impulses from sensory receptors to the CNS.
  2. **Motor or efferent neurons**: conduct impulses from the CNS to target organs (muscles or glands).
    - A. **Somatic motor neurons**: responsible for reflexes and voluntary control of skeletal muscles.
    - B. **Autonomic motor neurons**: innervate involuntary targets such as smooth muscle, cardiac muscle, and glands. **Two** subdivisions of autonomic neurons:
      - **Sympathetic** – emergency situations; “fight or flight”
      - **Parasympathetic** – normal functions; “rest and digest”
3. **Association/interneurons**: located completely within the CNS and integrate functions of the nervous system.

# Functional classification of Neurons





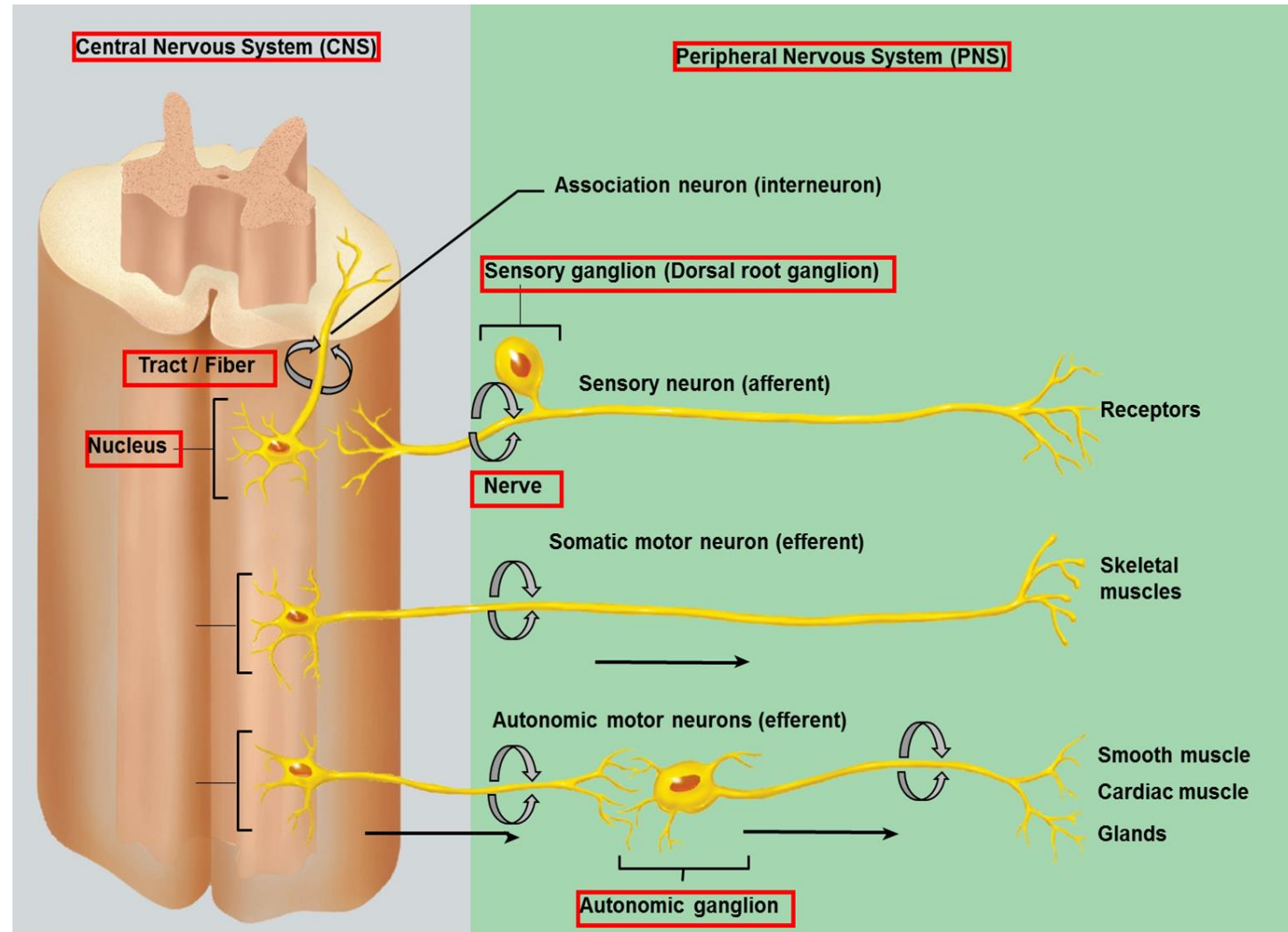
# Classification of Neurons and Nerves





# Classification of Nerves

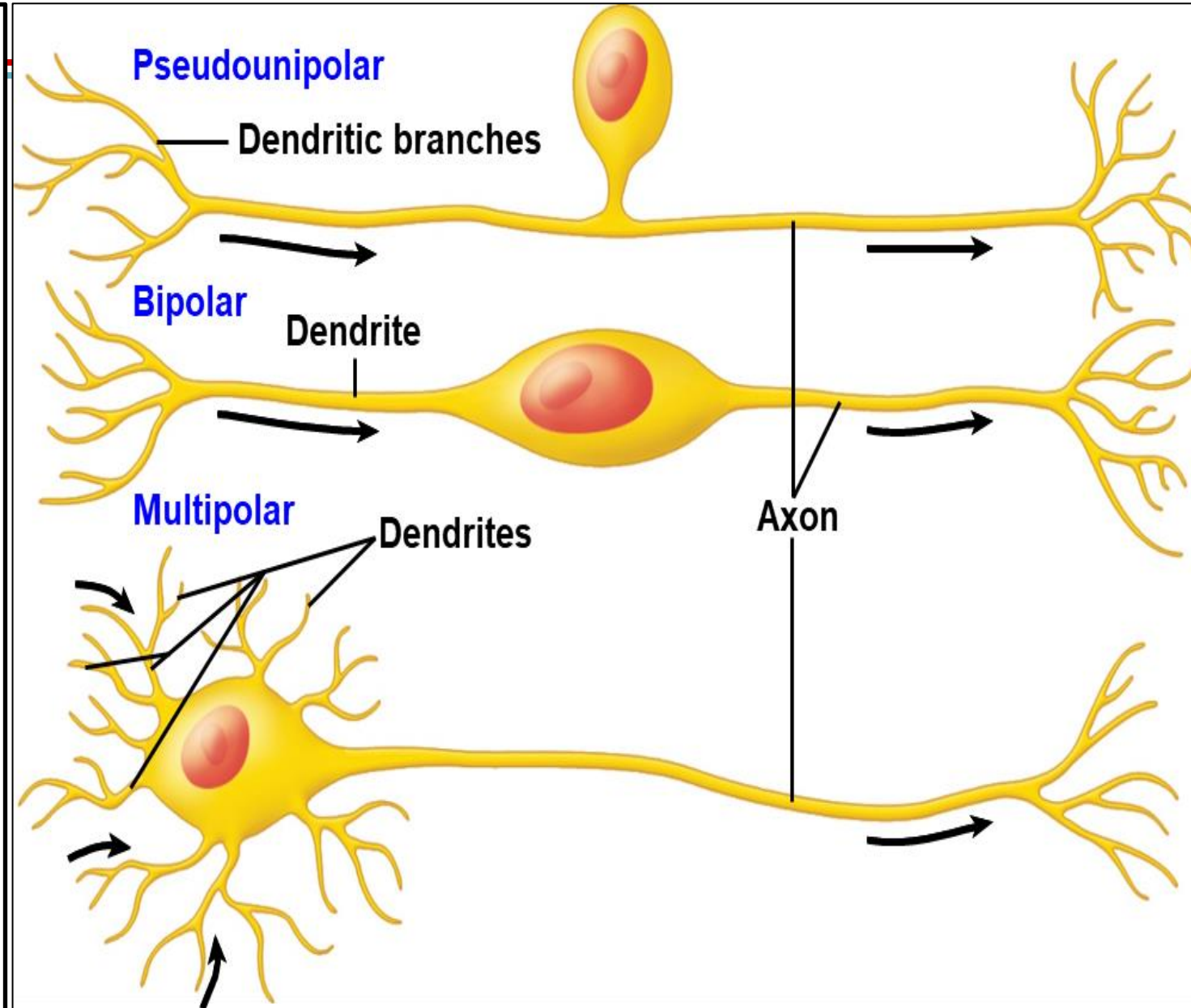
- Nerves are bundles of axons located outside the CNS.
- Most are composed of both sensory and motor neurons and are called mixed nerves.
- Some of the cranial nerves have sensory fibers only (nerves that serve the special senses of sight, hearing, and smell) .
- A bundle of axons in the CNS is called a tract.



# Structural Classification of Neurons

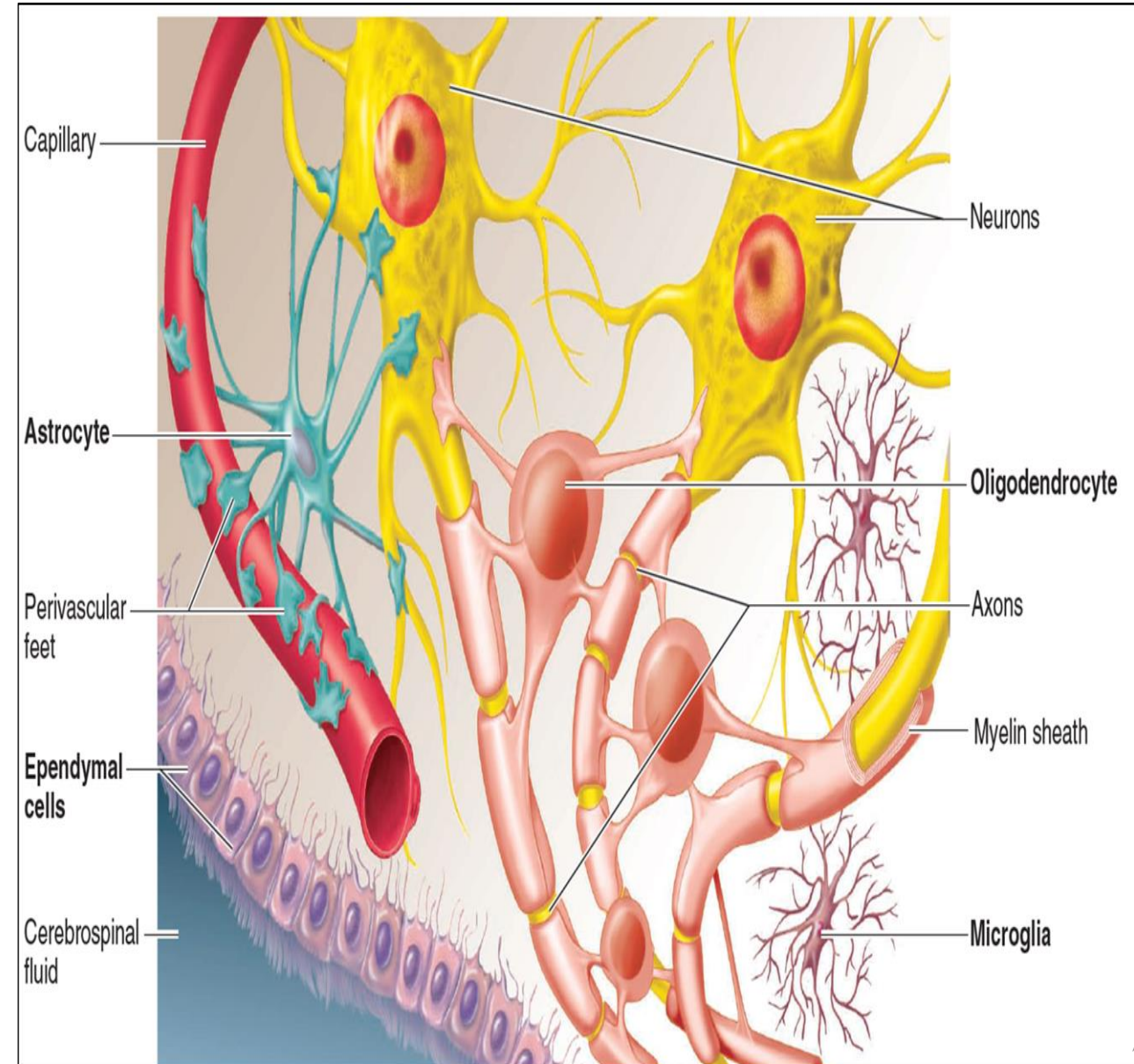
Based on the number of processes that extend from the cell body neurons are classified into:

- 1) **Pseudounipolar:** (from the Late Latin *pseudo* = false) single short process that branches like a T to form 2 longer processes; sensory neurons
- 2) **Bipolar neurons:** have two processes, one on either end; found in retina of eye
- 3) **Multipolar neurons:** several dendrites and one axon; most common type



# Neuroglia (glial cells)

- Cells that are non-conducting but support neurons
- Two types are found in the PNS:
  - 1) **Schwann cells (neurolemmocytes):** form myelin sheaths around peripheral axons
  - 2) **Satellite cells (ganglionic gliocytes):** support cell bodies within the ganglia of the PNS
- Four types are found in the CNS:
  - 1) **Oligodendrocytes:** form myelin sheaths around the axons of CNS neurons
  - 2) **Microglia:** migrate around CNS tissue and phagocytize foreign and degenerated material
  - 3) **Astrocytes:** regulate the external environment of the neurons
  - 4) **Ependymal cells:** line the ventricles and secrete cerebrospinal fluid





# Electrical Activity in Axons: Resting Membrane Potential

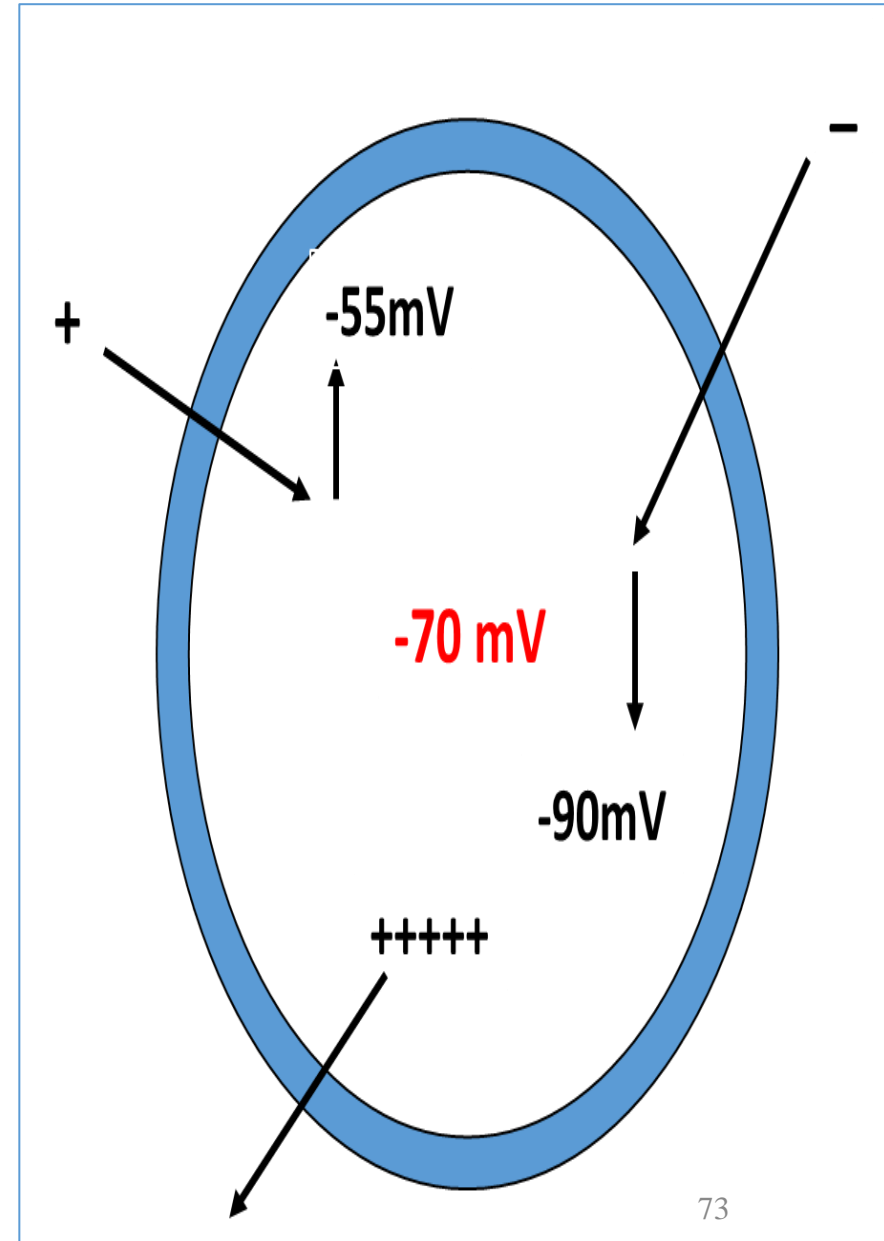
- Neurons have a resting potential of  $-70\text{mV}$ .
  - a. Established by large negative molecules inside the cell
  - b.  $\text{Na}^+/\text{K}^+$  pumps
  - c. Permeability of the membrane to positively charged, inorganic ions
- At rest, there is a high concentration of  $\text{K}^+$  inside the cell and  $\text{Na}^+$  outside the cell.

# Altering Membrane Potential

- A. Neurons and muscle cells can change their membrane potentials in response to **stimulation**.
- B. Called **excitability** or irritability
- C. Caused by **changes in the permeability** to certain ions
- D. Ions will follow their **electrochemical gradient** = combination of concentration gradient and attraction to opposite charges.
- E. Flow of ions are called **ion currents** which occur in limited areas where ion channels are located

## Changes in Membrane Potential

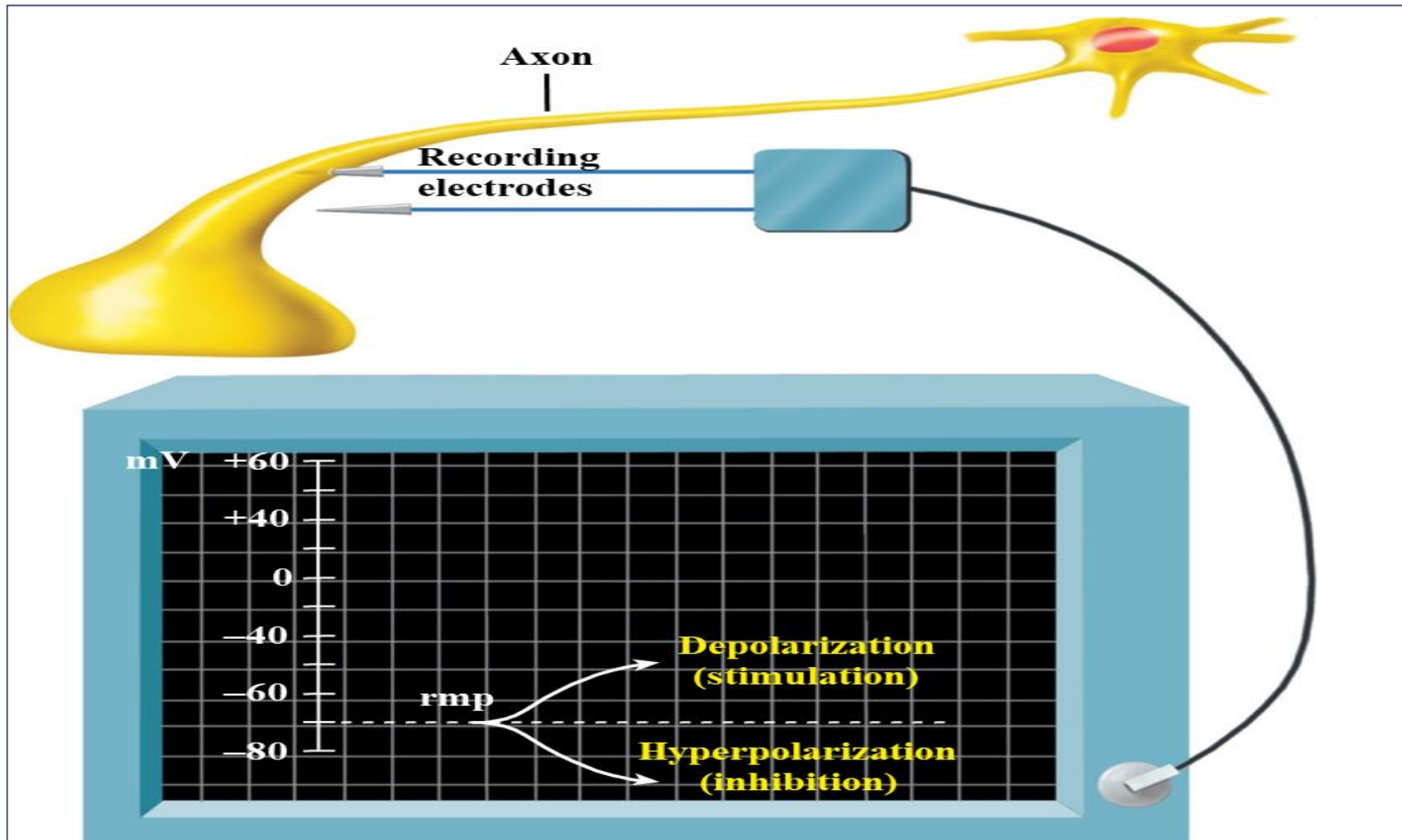
- At rest, a neuron is considered **polarized** when the inside is more negative than the outside.
- When the membrane potential inside the cell increases (becomes more positive), this is called **depolarization**.
- A return to resting potential is called **repolarization**.
- When the membrane potential inside the cell decreases (becomes more negative), this is called **hyperpolarization**.
- Depolarization occurs when positive ions enter the cell (usually  $\text{Na}^+$ ).
- Hyperpolarization occurs when positive ions leave the cell (usually  $\text{K}^+$ ) or negative ions ( $\text{Cl}^-$ ) enter the cell.
- Depolarization of the cell is **excitatory**.
- Hyperpolarization is **inhibitory**.





# Changes in Membrane Potential

- Changes can be recorded on an **Oscilloscope** by recording the voltage inside and outside the cell.



# Ion Gating in Axons

- **Changes in membrane potential are controlled** by changes in the flow of ions through channels.

A.  $K^+$  has two types of channels:

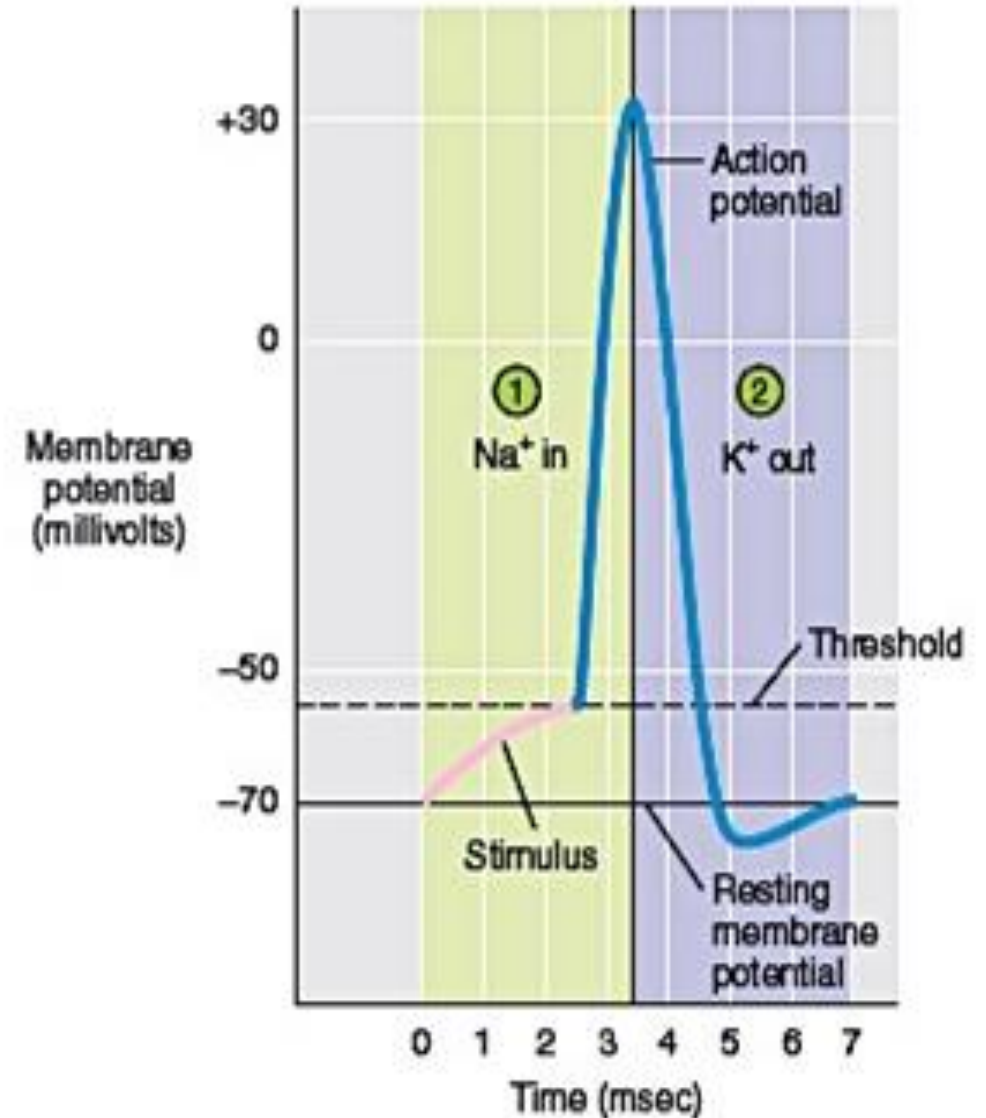
1). **Not gated** (always open); sometimes called  $K^+$  leakage channels.

2). **Voltage-gated  $K^+$  channels**; open when a particular membrane potential is reached; closed at resting potential

B.  $Na^+$  has only channels that are closed at rest; the membrane is less permeable to  $Na^+$  at rest (**Voltage-gated  $Na^+$  channels**).

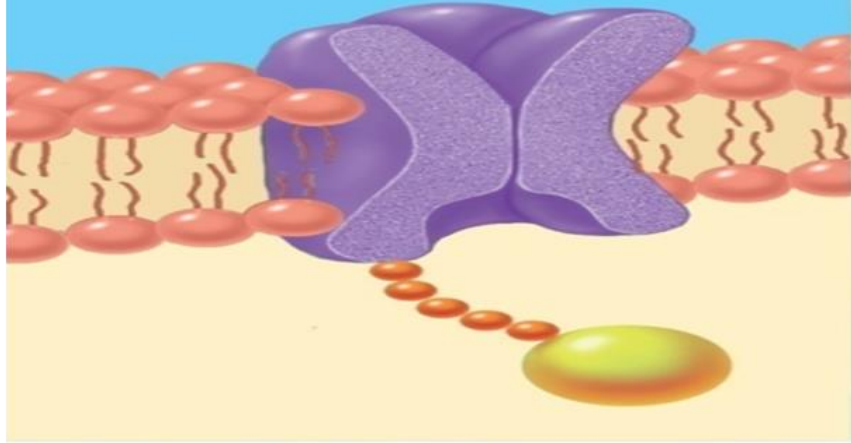
# Voltage-Gated $\text{Na}^+$ Channels

- a. At rest, the  $\text{Na}^+$  channels are **closed**.
- b. These channels **open** if the membrane potential depolarizes to  **$-55\text{mV}$** .
- c. This is called the **threshold**.
- d. Sodium rushes in due to the electrochemical gradient.
- e. Membrane potential climbs toward sodium equilibrium potential.
- f. These channels become **inactivated** at  $+30\text{mV}$ .

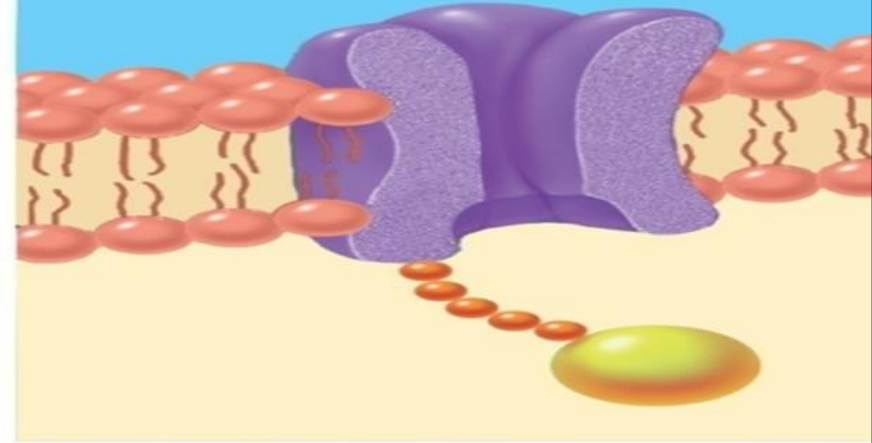


# A Voltage-Gated Ion Channel

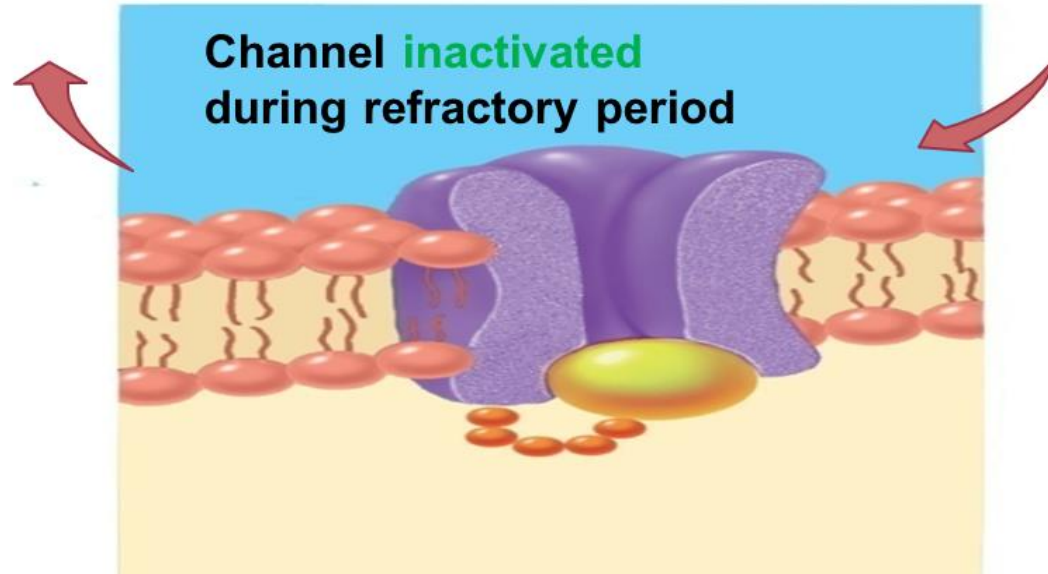
Channel **closed** at resting membrane potential



Channel **open** by depolarization (action potential)



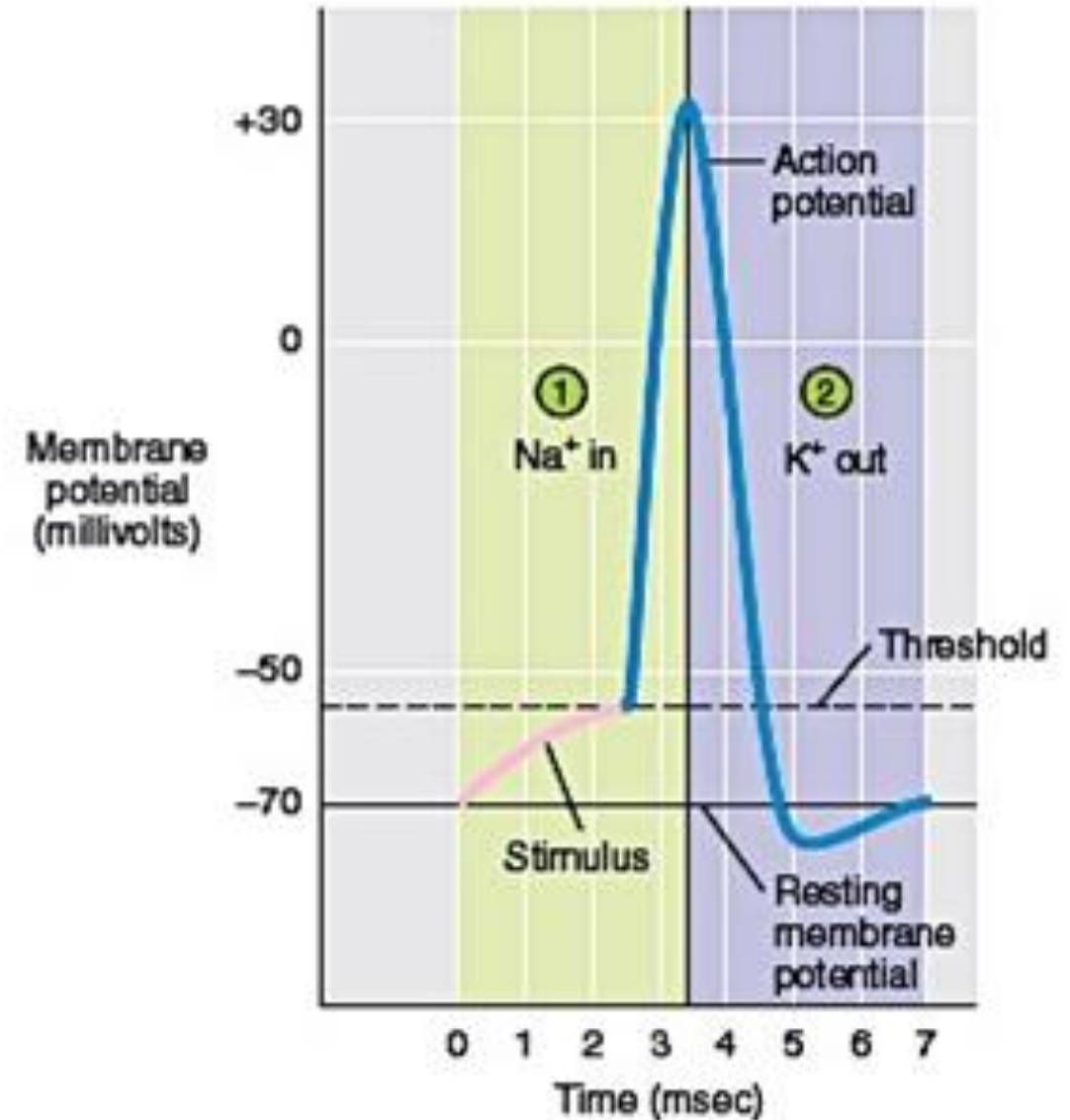
Channel **inactivated** during refractory period



# Voltage-Gated $K^+$ Channels

A. At around +30 mV, voltage-gated  $K^+$  channels open, and  $K^+$  rushes out of the cell following the electrochemical gradient.

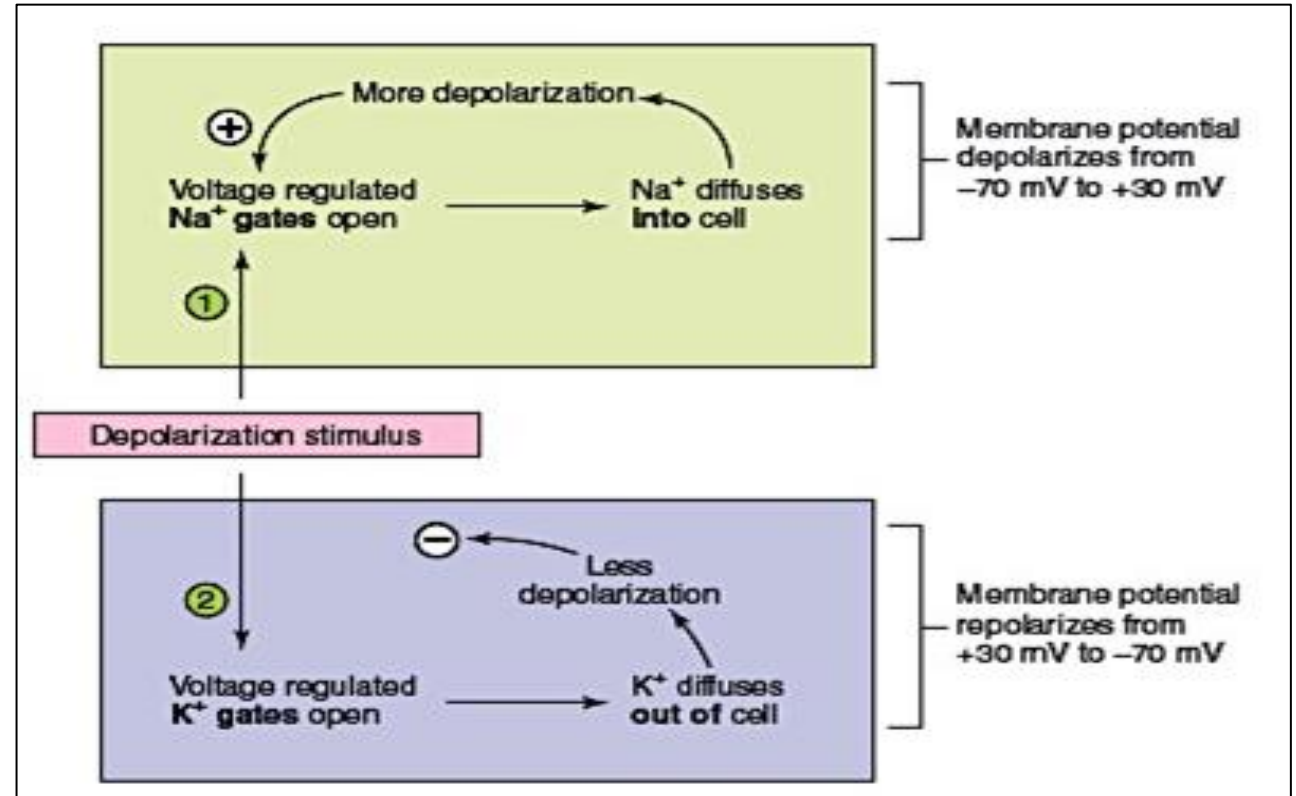
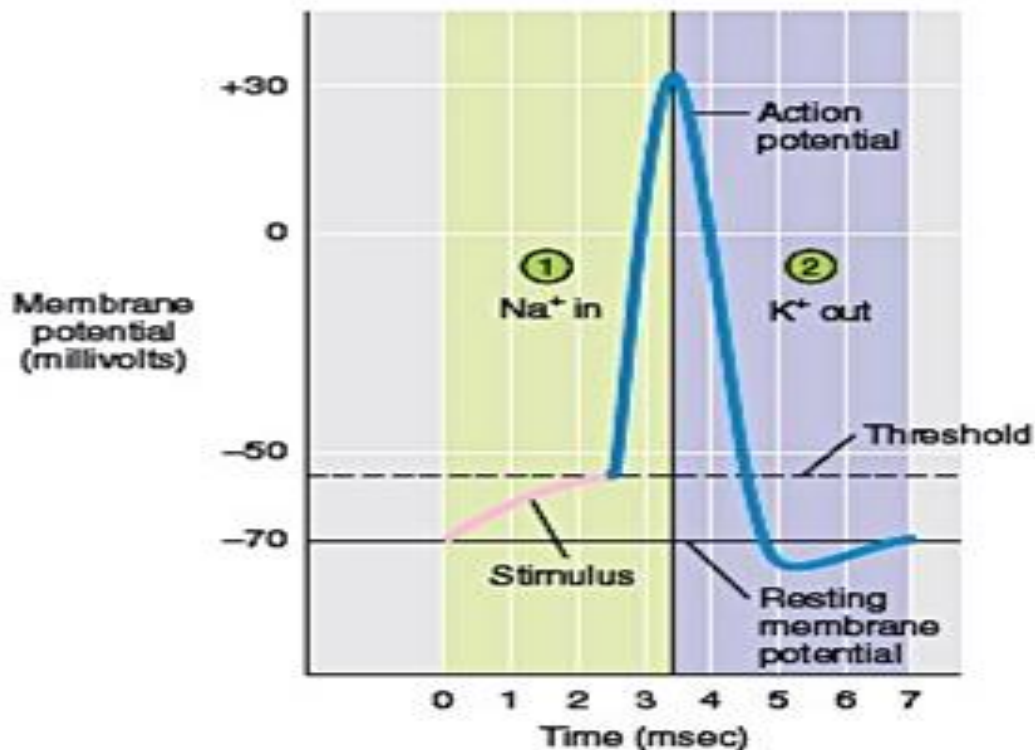
B. This makes the cell repolarize back toward the potassium equilibrium potential.





# Action Potentials (nerve impulse)

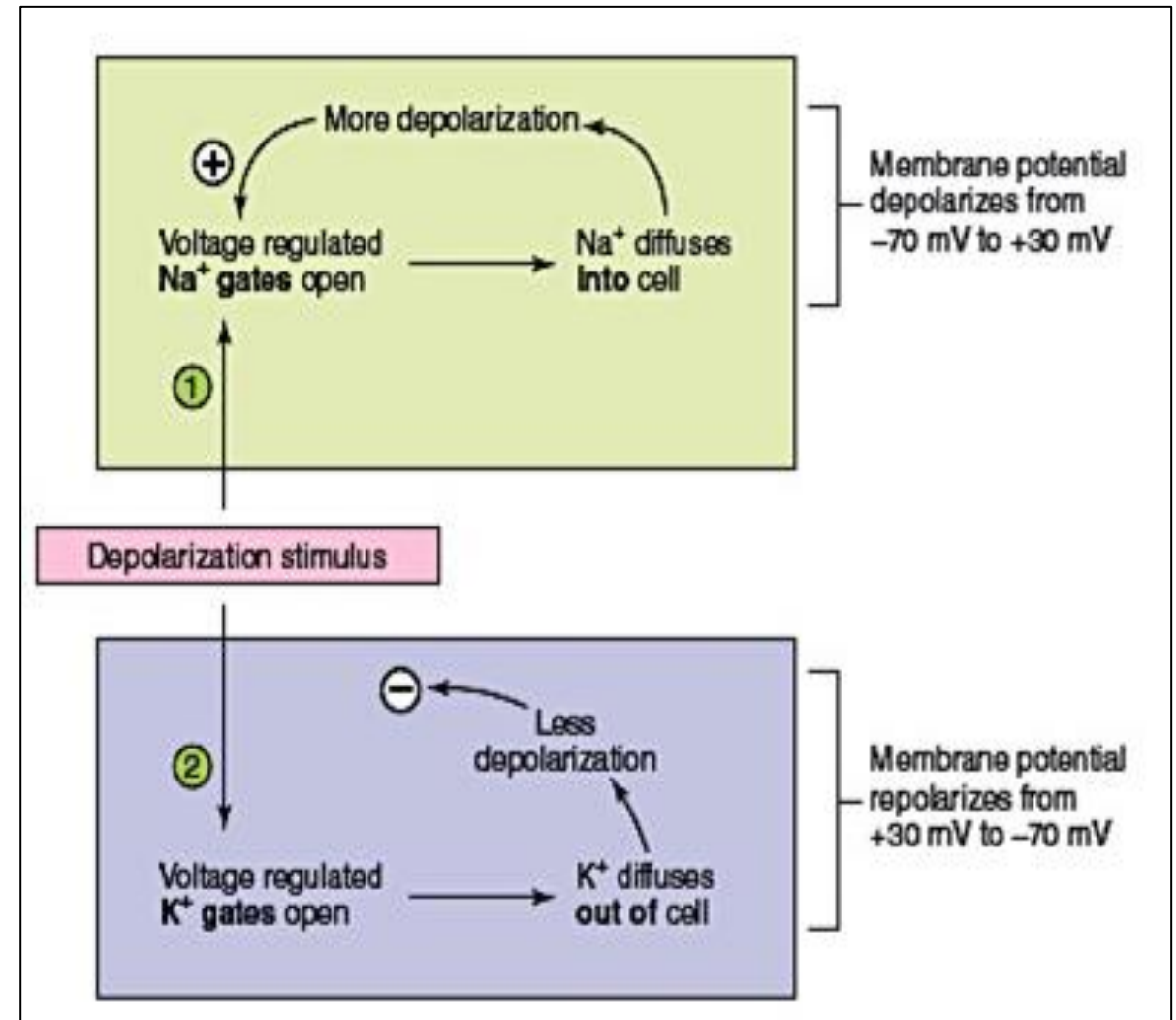
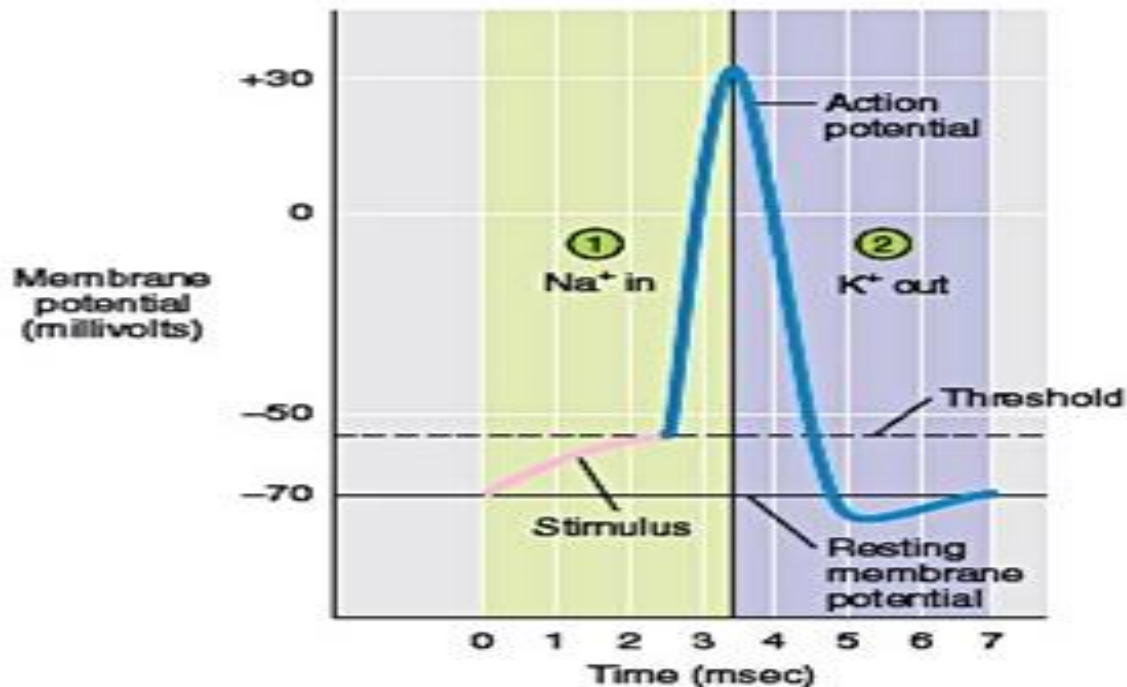
- At threshold membrane potential ( $-55\text{mV}$ ), voltage-gated  $\text{Na}^+$  channels open, and  $\text{Na}^+$  rushes in. As the cell depolarizes, more  $\text{Na}^+$  channels are open, and the cell becomes more and more permeable to  $\text{Na}^+$ .
  - This is a **positive feedback loop**.
  - Rapidly depolarize to  $0\text{mV}$  and then *overshoot* of the membrane potential to reach  $+30\text{mV}$  (depolarization)





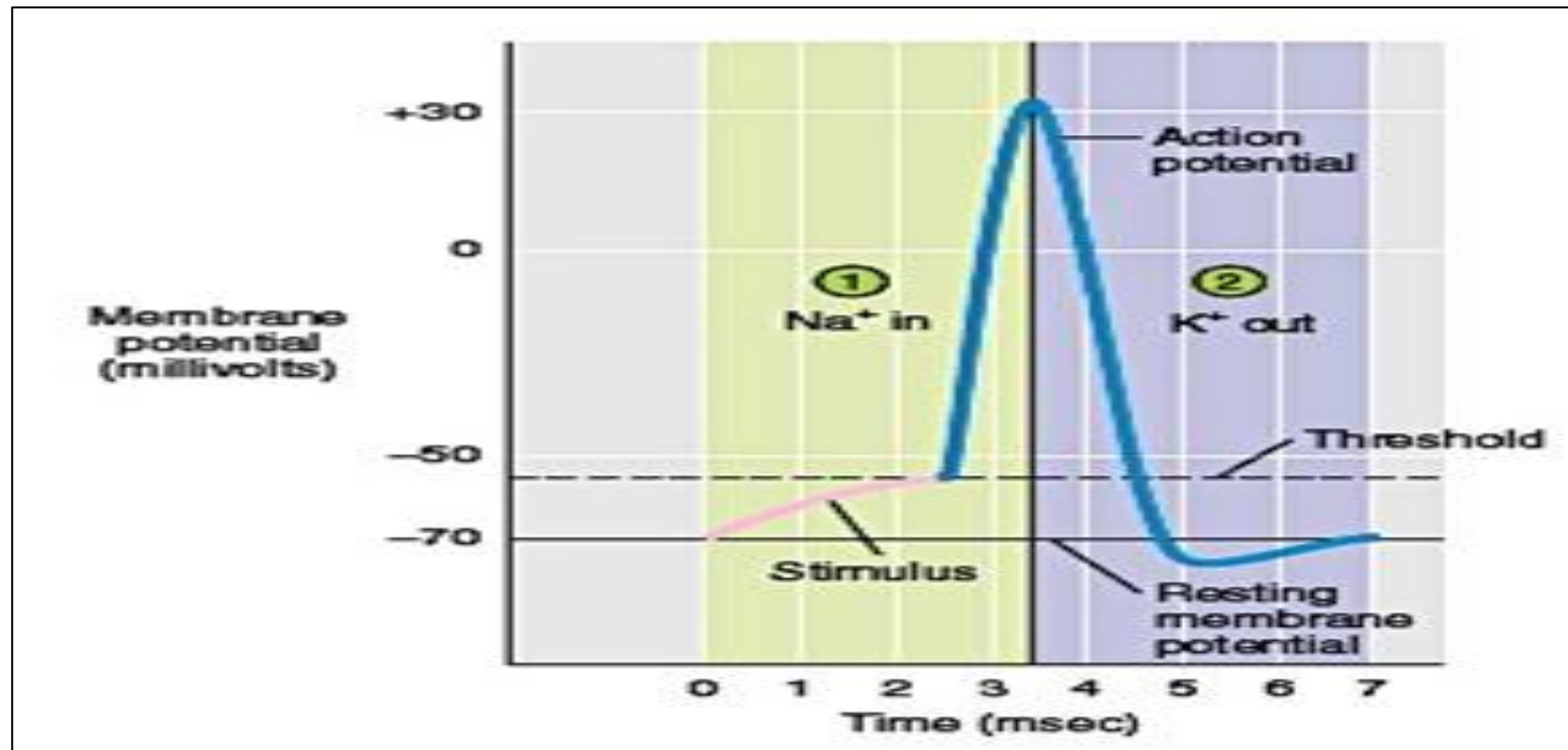
# Action Potentials (nerve impulse)

2. At +30mV,  $\text{Na}^+$  channels become inactivated, and  $\text{K}^+$  channels open.
- Results in repolarization of membrane potential
  - This is a **negative feedback loop**



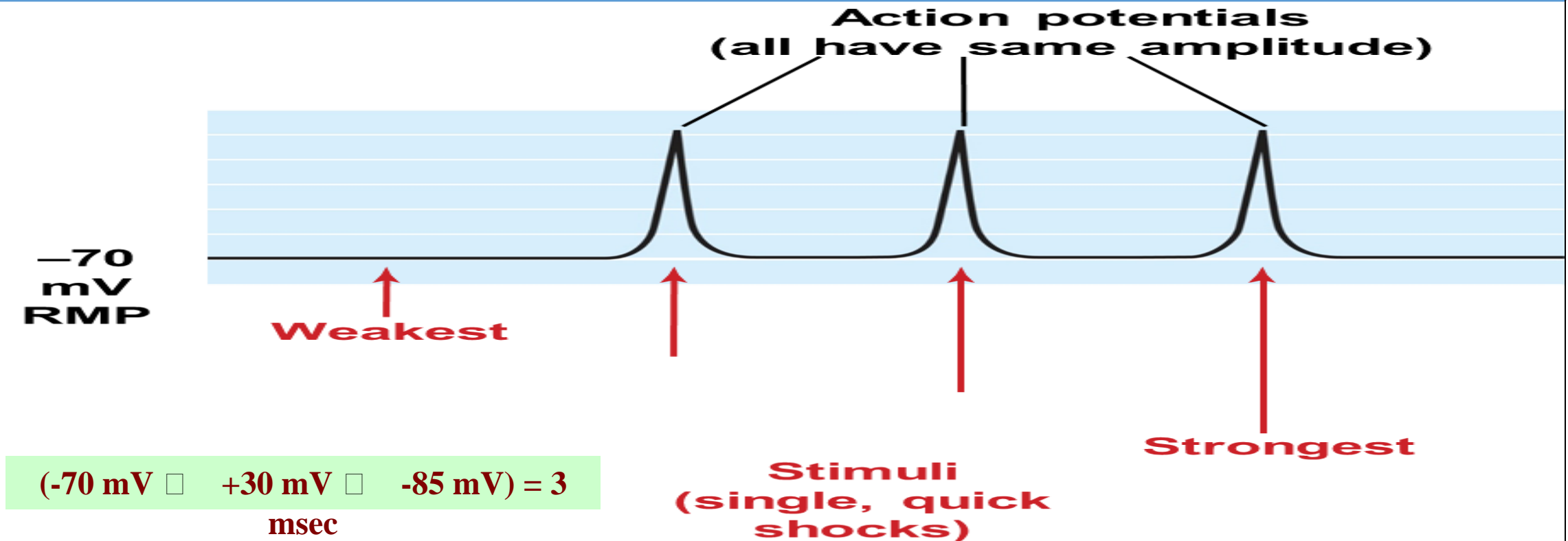
## After-Hyperpolarization

- a. Repolarization actually overshoots resting potential and gets down to  $-85\text{mV}$  as a result of **continued outward movement of  $\text{K}^+$** .
- b. This does not reach potassium equilibrium potential because voltage-gated  **$\text{K}^+$  channels are inactivated** as the membrane potential falls.
- c.  $\text{Na}^+/\text{K}^+$  pumps **quickly reestablish resting potential**.



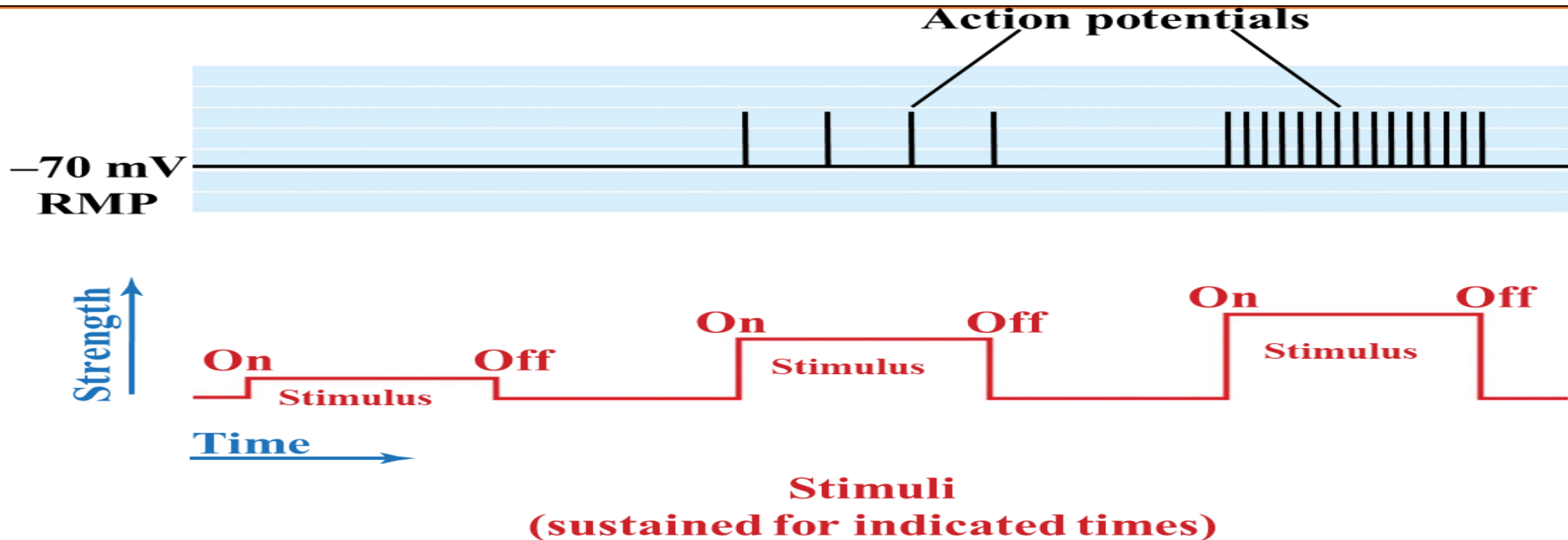
# All-or-None Law

- a. Once threshold has been reached, an action potential will happen.
- b. The size of the stimulus will not affect the size of the action potential; it will always reach +30mV.
- c. The size of the stimulus will not affect action potential duration.



## Coding for Stimulus Intensity (the effect of stimulus strength on action potential frequency)

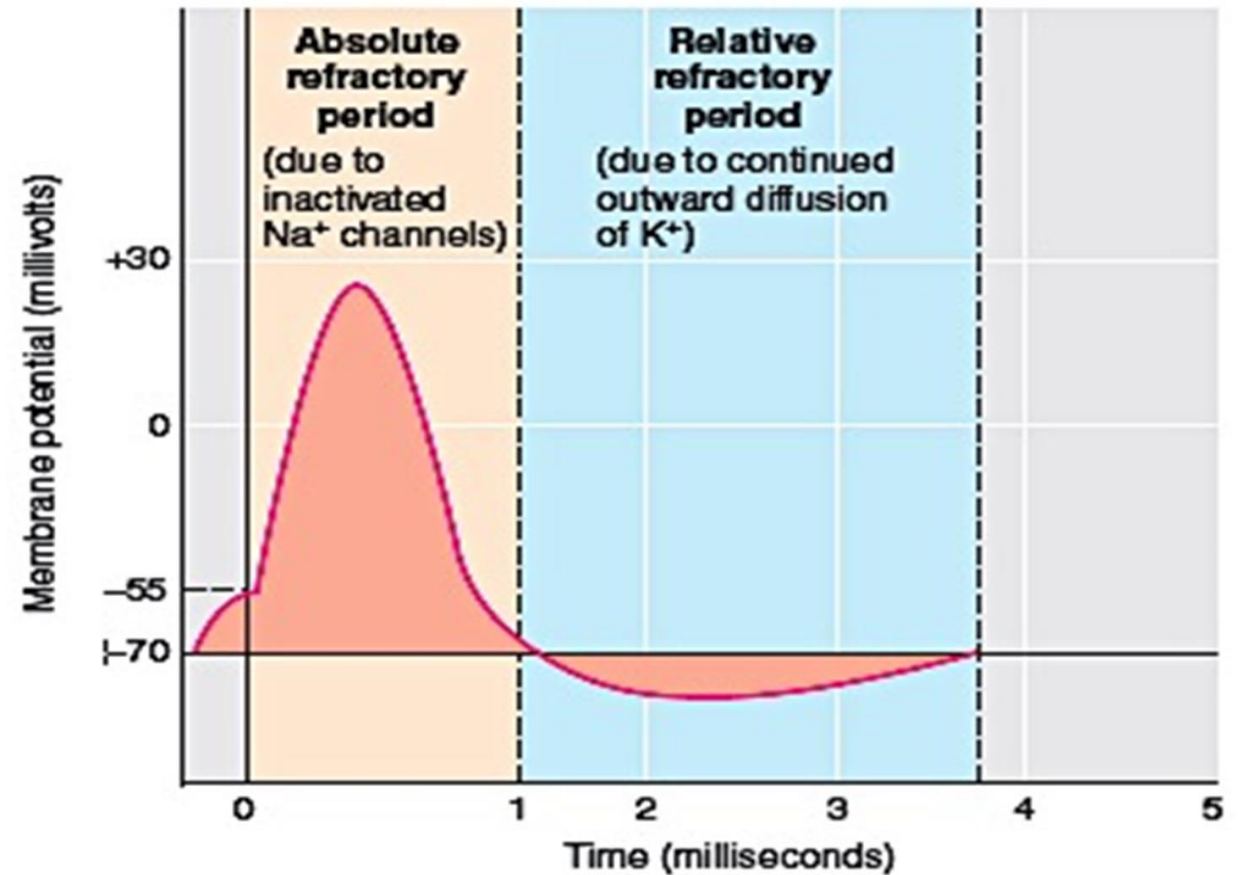
- A stronger stimulus will make identical action potentials occur more frequently (**frequency modulated**)
- A stronger stimulus may also activate more axons in a nerve (**recruitment**)



Stimulus strength is coded by the frequency rather than the amplitude of action potentials

# Refractory Periods

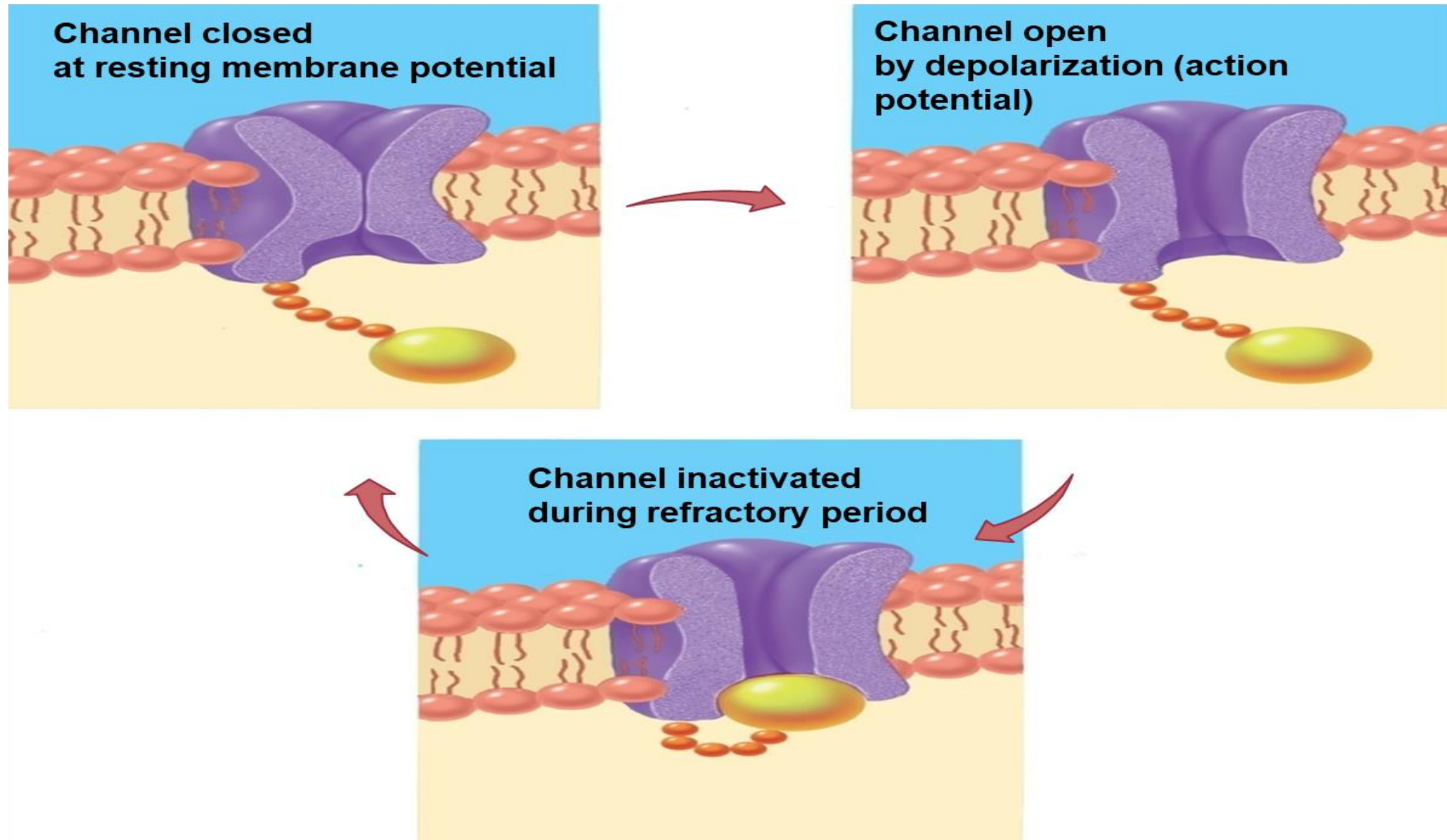
- a. Action potentials can only increase in frequency to a certain point. There is a refractory period after an action potential when the neuron cannot become excited again.
- b. The **absolute refractory period** occurs during the action potential. **Na<sup>+</sup> channels are inactive (not just closed).**
- c. The **relative refractory period** is when K<sup>+</sup> channels are still open. Only a very strong stimulus can overcome this.
- d. Each action potential remains a separate, all-or-none event.



**Figure 7.17 Absolute and relative refractory periods.** While a segment of axon is producing an action potential, the membrane is absolutely or relatively resistant (refractory) to further stimulation. **AP|R**



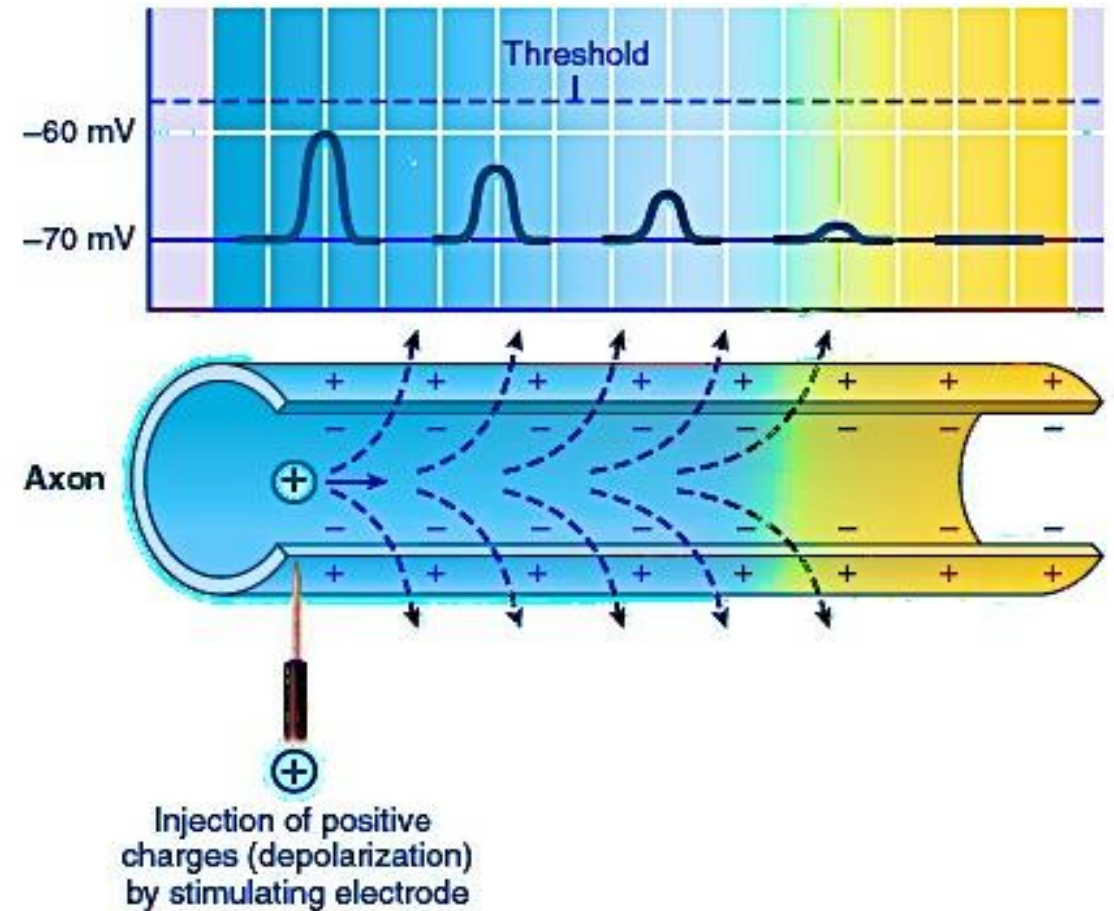
# A Voltage-Gated Ion Channel





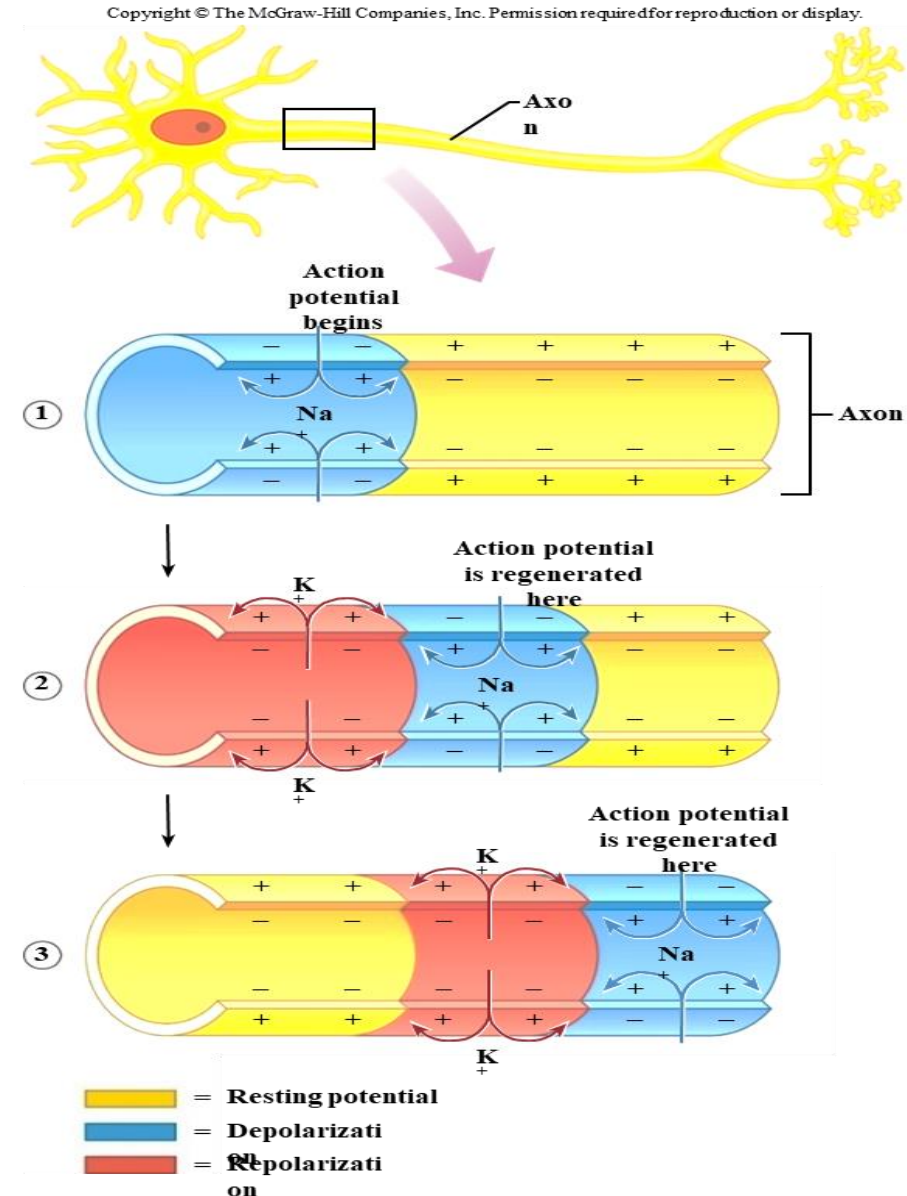
# Cable Properties of Neurons

- a. The ability of neurons to conduct charges through their cytoplasm
- b. Poor due to high internal resistance to the spread of charges and leaking of charges through the membrane
- c. Neurons could not depend on cable properties to move an impulse down the length of an axon.



# Conduction of Nerve Impulses

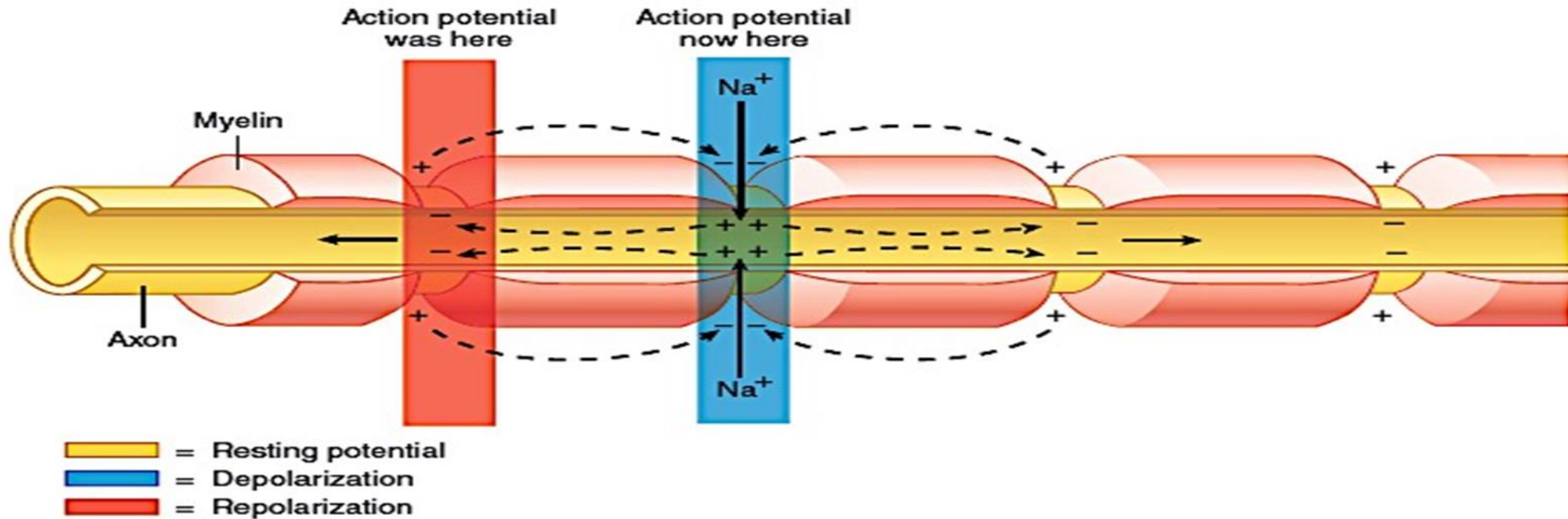
1. When an action potential occurs at a given point on a neuron membrane, voltage-gated  $\text{Na}^+$  channels open as a wave down the length of the axon.
2. The action potential at one location serves as the depolarization stimulus for the next region of the axon



## Conduction in an **unmyelinated** neuron

- a. Axon potentials are **produced down the entire length of the axon** at every patch of membrane.
- b. The conduction rate is **slow** because so many action potentials are generated, each one, an individual event.
- c. The **amplitude** of each action potential is the **same** — conducted without decrement.

# Conduction in a myelinated neuron



- Myelin provides insulation, improving the speed of cable properties.
- Nodes of Ranvier allow  $\text{Na}^+$  and  $\text{K}^+$  to cross the membrane every 1–2 mm.
- $\text{Na}^+$  ion channels are concentrated at the nodes
- Action potentials “leap” from node to node
- This is called saltatory conduction.

# Action Potential Conduction Speed

## a. Increased by:

- 1) Increased diameter of the neuron. This reduces resistance to the spread of charges via cable properties.
- 2) Myelination because of saltatory conduction

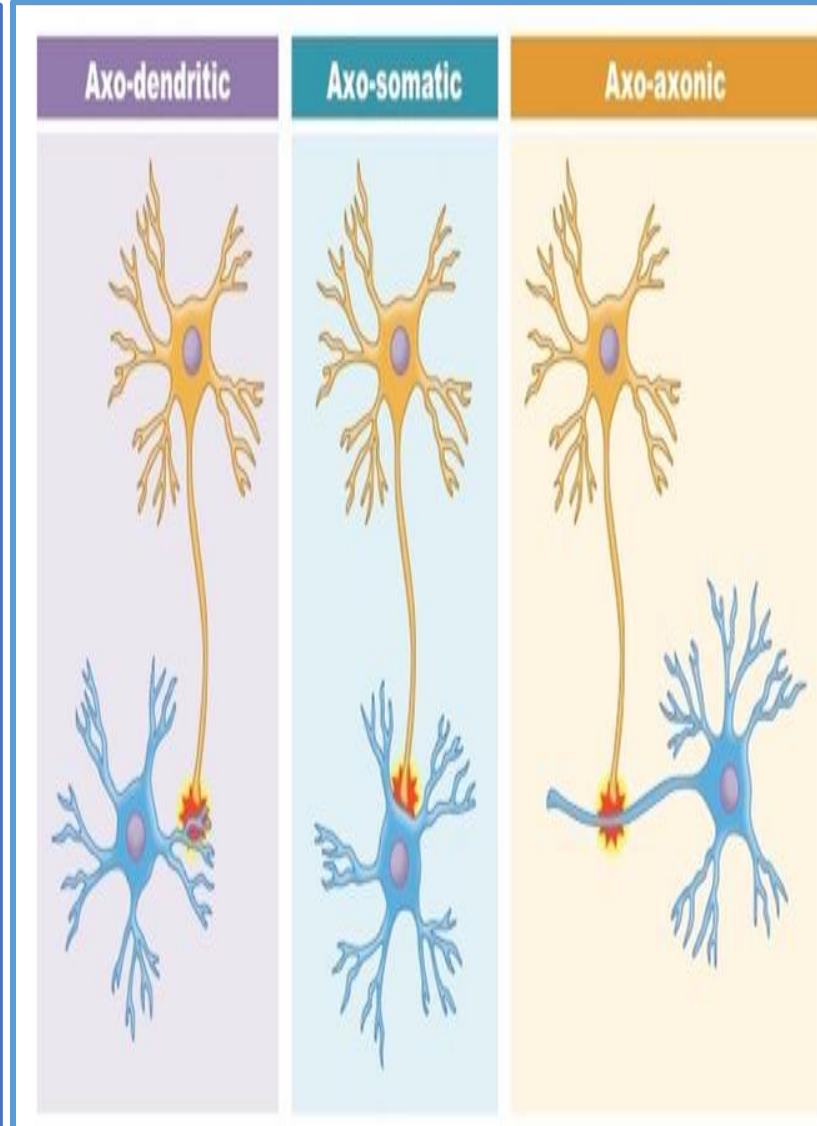
## b. Examples

- 1) Thin, unmyelinated neuron speed 1.0m/sec
- 2) Thick, myelinated neuron speed 100m/sec



# The Synapse

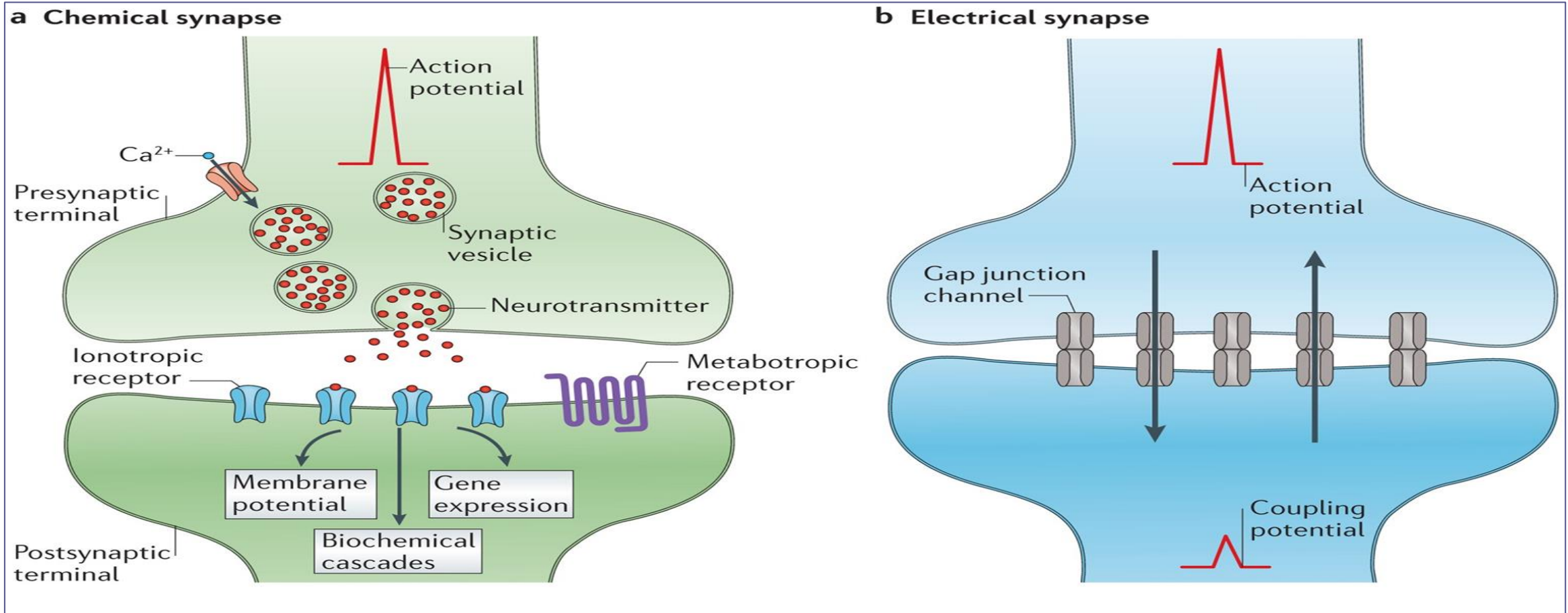
- A synapse is the functional connection between a neuron and the cell it is signaling
  - a. In the **CNS**, this second cell will be another neuron.
  - b. In the **PNS**, the second cell will be in a muscle or gland; often called myoneural or neuromuscular junctions
- 2. If one neuron is signaling another neuron, the first is called the **presynaptic** neuron, and the second is called the **postsynaptic** neuron.
  - A presynaptic neuron can signal the dendrite, cell body, or axon of a second neuron.
  - There are **axodendritic**, **axosomatic**, and **axoaxonic** synapses. **Axosomatic synapses** are synapses that are made onto the soma or cell body of a neuron. **Axodendritic synapses**, probably the most prominent kind of synapses, are synapses that one neuron makes onto the dendrite of another neuron. **Axoaxonic synapses** are synapses made by one neuron onto the synapse of another neuron.
  - Most synapses are axodendritic and are one direction
- 3. Synapses can be **electrical** or **chemical**





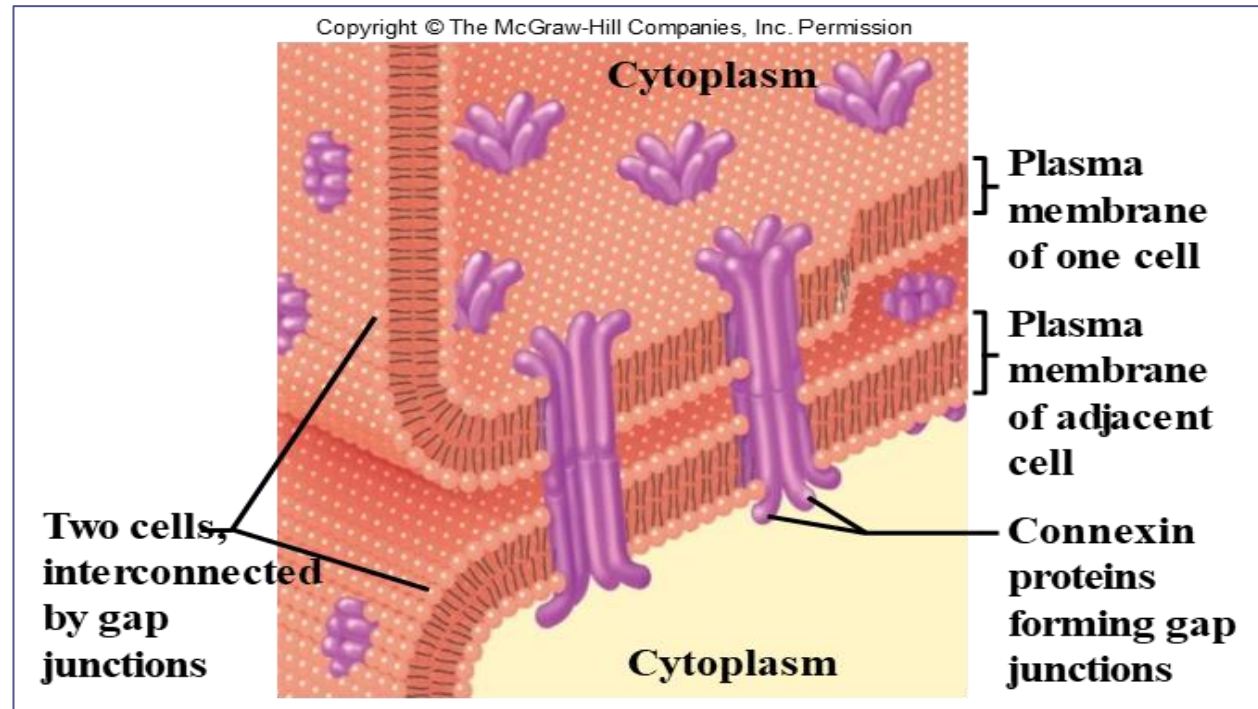
## Electrical Synapses versus Chemical Synapses

- **Electrical synapses** occur in smooth muscle and cardiac muscle, between some neurons of the brain, and between glial cells. Cells are joined by **gap junctions**.
- **Chemical synapses** involve the release of a chemical called a **neurotransmitter** from the axon's terminal boutons into a very small **synaptic cleft** between the **presynaptic cell** and **postsynaptic cell**.



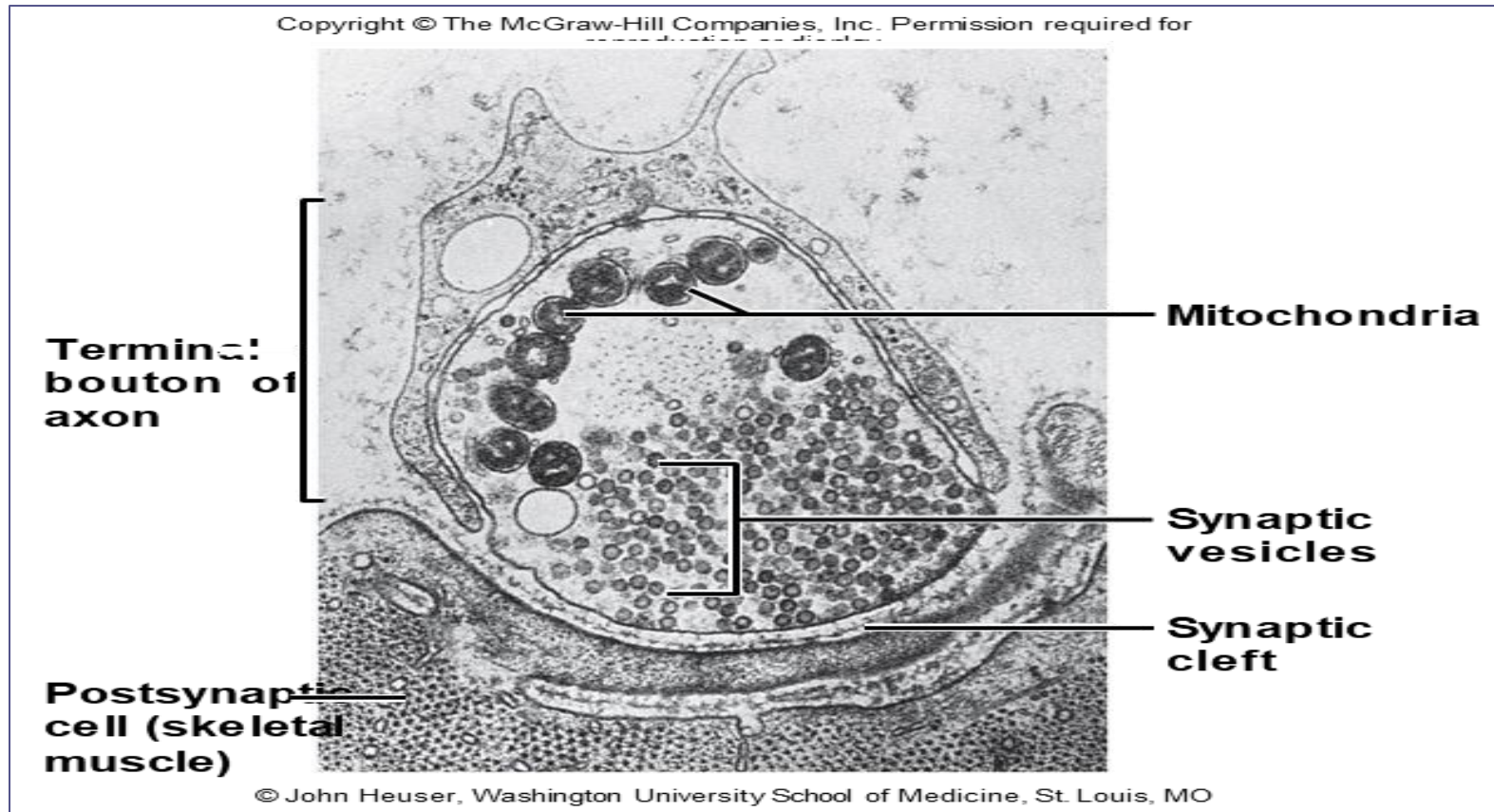
# Electrical Synapses

- Cells are joined by gap junctions. In gap junctions, the membranes of the two cells are separated by only 2 nanometers. Ions and molecules can pass from one cell to the other.
- Stimulation causes phosphorylation or dephosphorylation of **connexin** proteins to open or close the channels
- Electrical synapses occur in smooth muscle and cardiac muscle, between some neurons of the brain, and between glial cells.



# Chemical Synapses

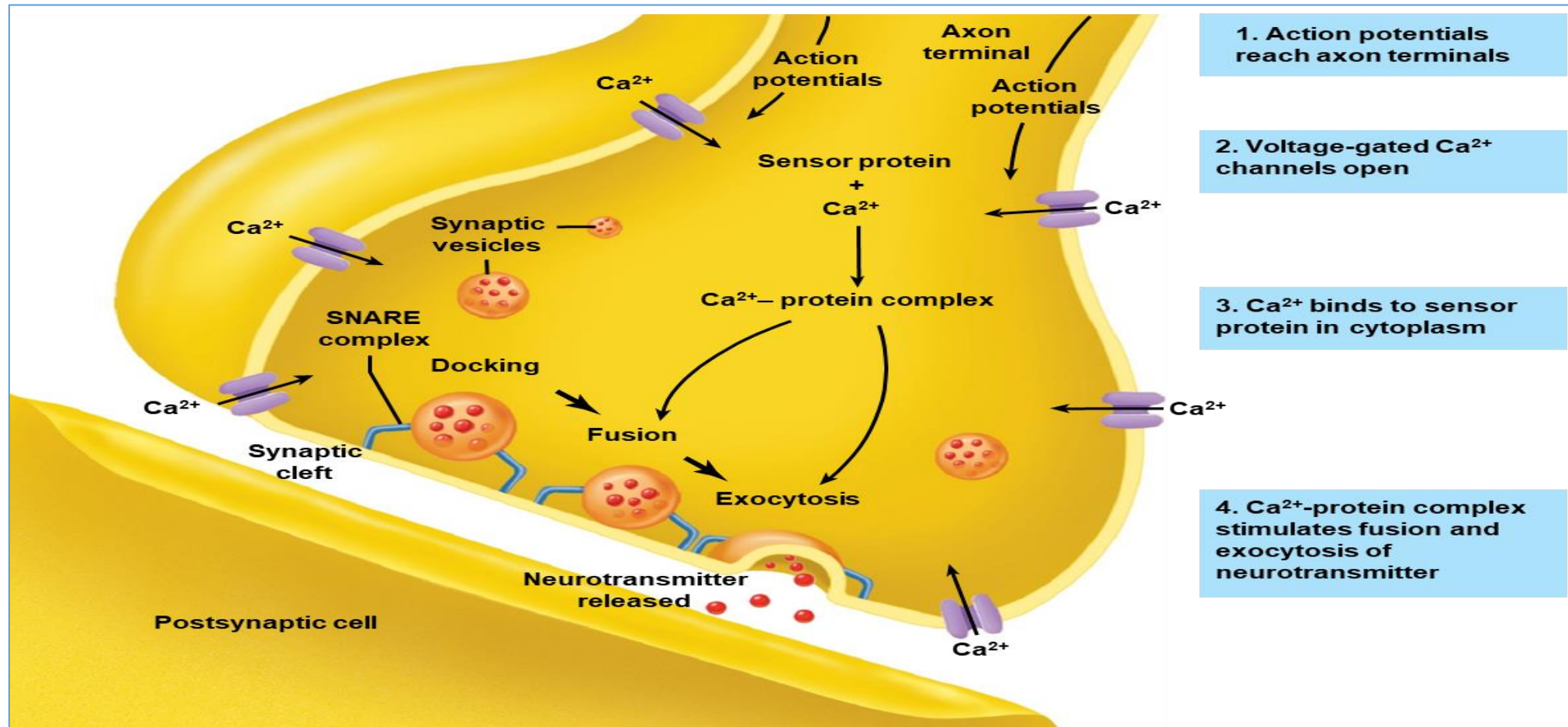
- Most synapses involve the release of a chemical called a neurotransmitter from the axon's terminal buttons.
- The synaptic cleft is very small, and the presynaptic and postsynaptic cells are held close together by cell adhesion molecules (CAMs).





# Release of Neurotransmitter

- Neurotransmitter is enclosed in synaptic vesicles in the axon terminal.
  - When the action potential reaches the end of the axon, **voltage-gated  $\text{Ca}^{2+}$  channels** open.
  - $\text{Ca}^{2+}$  stimulates the fusing of synaptic vesicles to the plasma membrane and **exocytosis** of neurotransmitter.**
  - A greater frequency** of action potential results in more stimulation of the **postsynaptic** neuron.



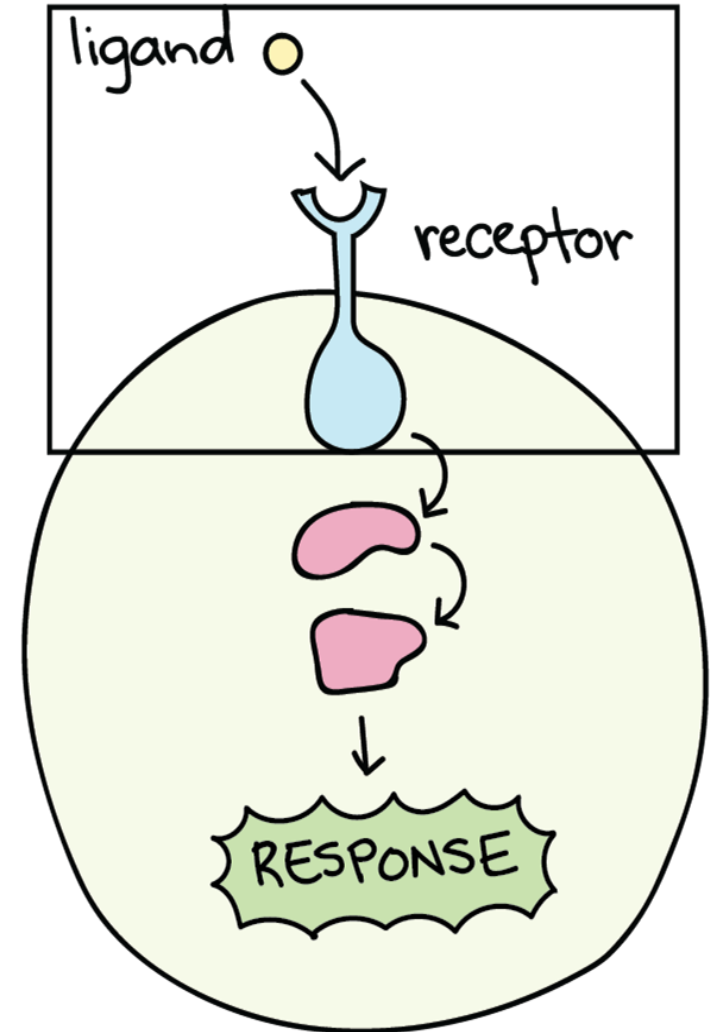
## Ca<sup>2+</sup> and Synaptic Vesicles

- 1) When Ca<sup>2+</sup> enters the cell, it binds to a protein called **synaptotagmin** that serves as a Ca<sup>2+</sup> sensor
- 2) Vesicles containing neurotransmitter are docked at the plasma membrane by three **SNARE proteins**.
- 3) The **Ca<sup>2+</sup> synaptotagmin** complex stimulates vesicle fusion to the membrane
- 4) Forms a pore to release the NT

# Action of a Neurotransmitter

- The neurotransmitter diffuses across the synapse, where it binds to a **specific receptor protein on the postsynaptic cell**.

- 1) The neurotransmitter is referred to as the **ligand**.
- 2) Binding of the neurotransmitter to the receptor results in the opening/closing of ion channels (so-called chemically regulated ion channels) on the plasma membrane of the postsynaptic cell **or** activates second messenger systems inside the postsynaptic cell.





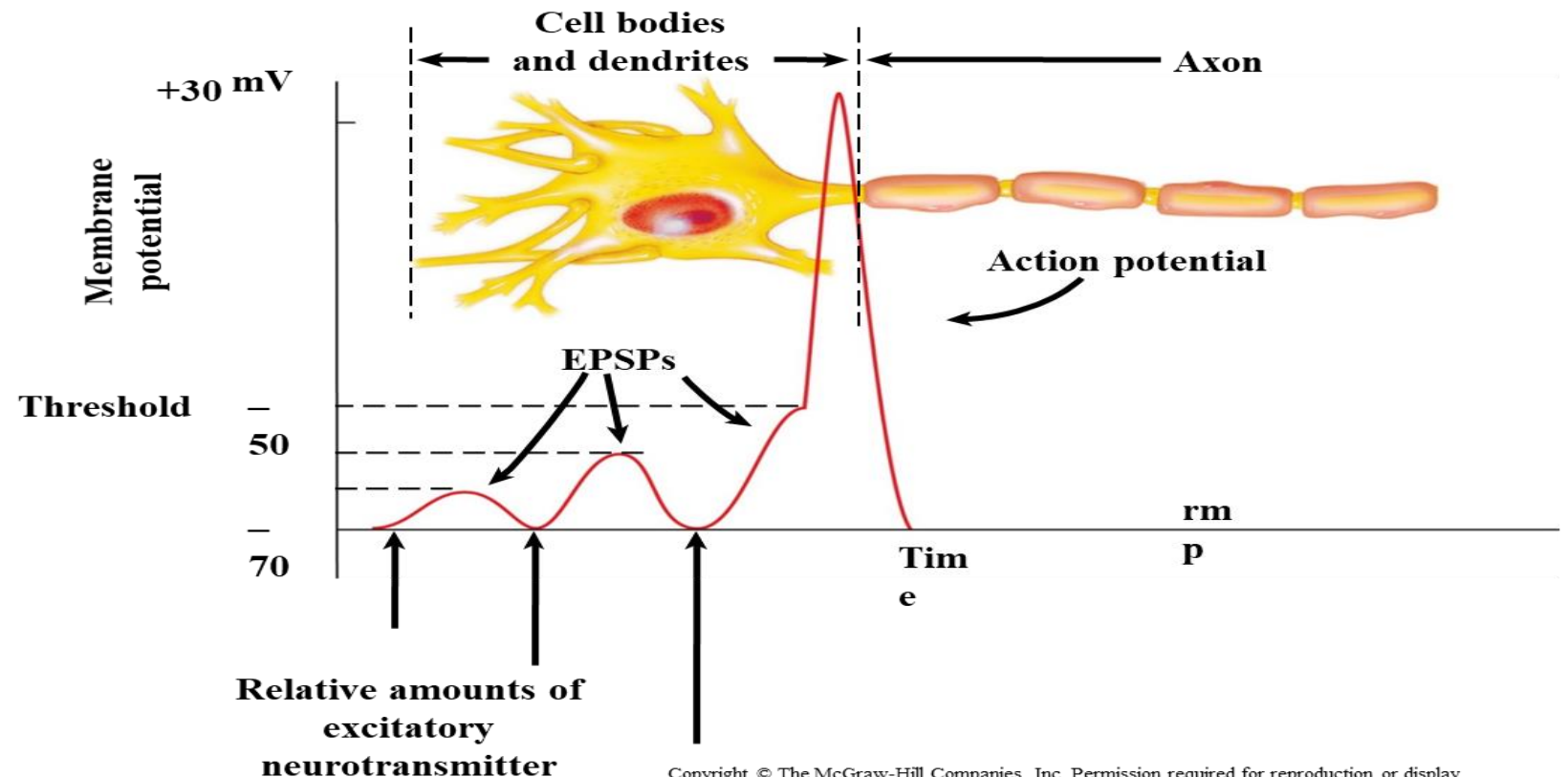
# Graded Potential

- 1) Graded change in the membrane potential of the chemically regulated ion channels.
- 2) When ligand-gated ion channels open, the membrane potential changes **depending on which ion channel is open**.
  - a) Opening  $\text{Na}^+$  or  $\text{Ca}^{2+}$  channels results in a graded depolarization called an **excitatory postsynaptic potential (EPSP)**.
  - b) Opening  $\text{K}^+$  or  $\text{Cl}^-$  channels results in a graded hyperpolarization called **inhibitory postsynaptic potential (IPSP)**.

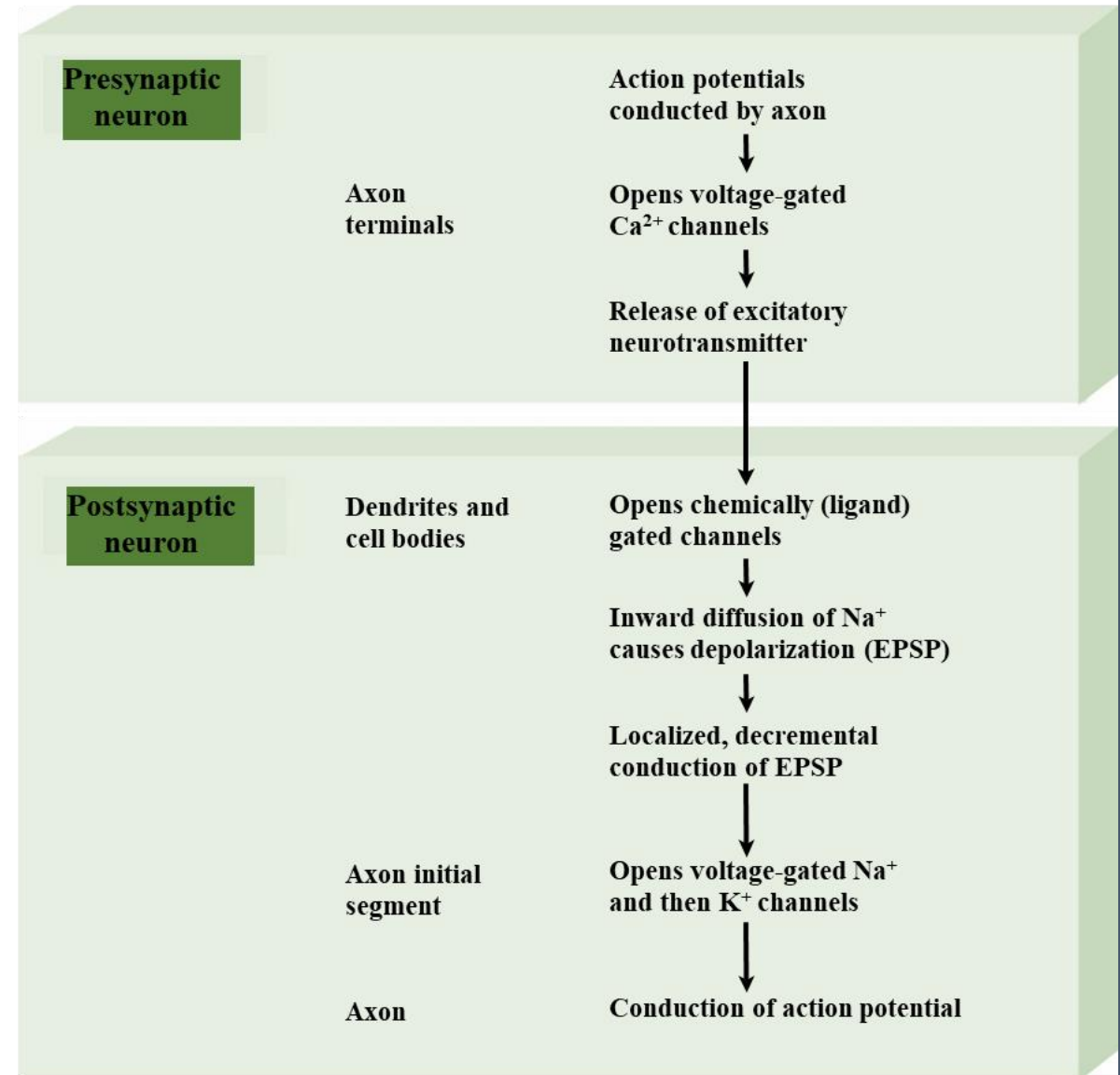
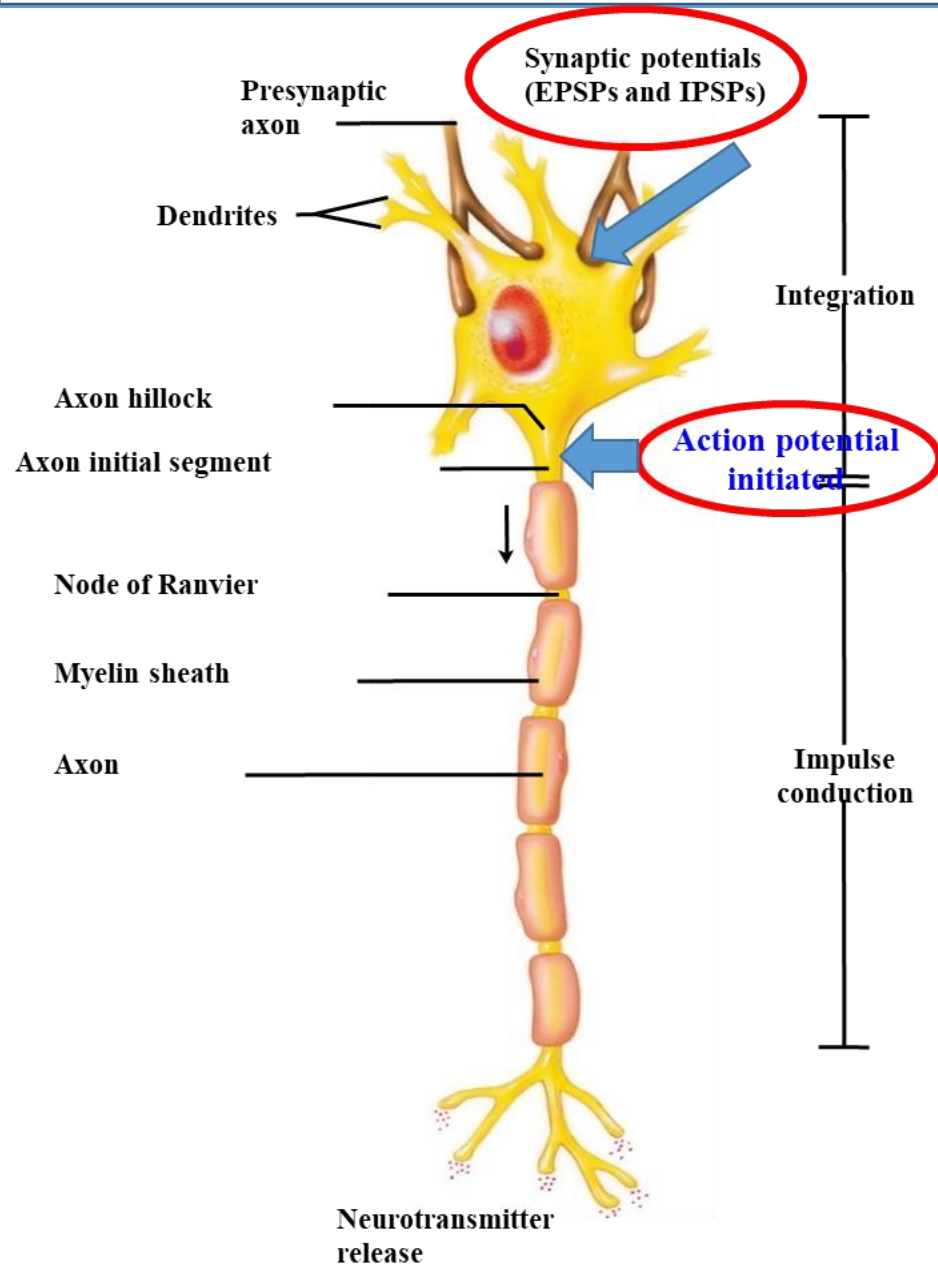
# EPSPs and IPSPs

- EPSPs move the membrane potential **closer** to threshold; may require EPSPs from several neurons to actually produce an action potential.
- IPSPs move the membrane potential **farther** from threshold.
- Can counter EPSPs from other neurons so summation of EPSPs and IPSPs at the initial segment of the axon (next to the axon hillock) determines whether an action potential occurs.

## Graded nature of EPSPs



# Summary of Neurotransmitter Action



# Acetylcholine (ACh)

- ACh is a neurotransmitter that directly opens ion channels when it binds to its receptor.
  - a. In some cases, ACh is **excitatory**, and in other cases it is **inhibitory**, depending on the organ involved.
  - b. **Excitatory** in some areas of the CNS, in some autonomic motor neurons, and in all somatic motor neurons
  - c. **Inhibitory** in some autonomic motor neurons

# Two types of acetylcholine receptors

## a. Nicotinic ACh receptors

- 1) Can be stimulated by nicotine
- 2) Found on the motor end plate of skeletal muscle cells, in autonomic ganglia, and in some parts of the CNS

## b. Muscarinic ACh receptors

- 1) Can be stimulated by muscarine (from poisonous mushrooms)
- 2) Found in CNS and plasma membrane of smooth and cardiac muscles and glands innervated by autonomic motor neurons

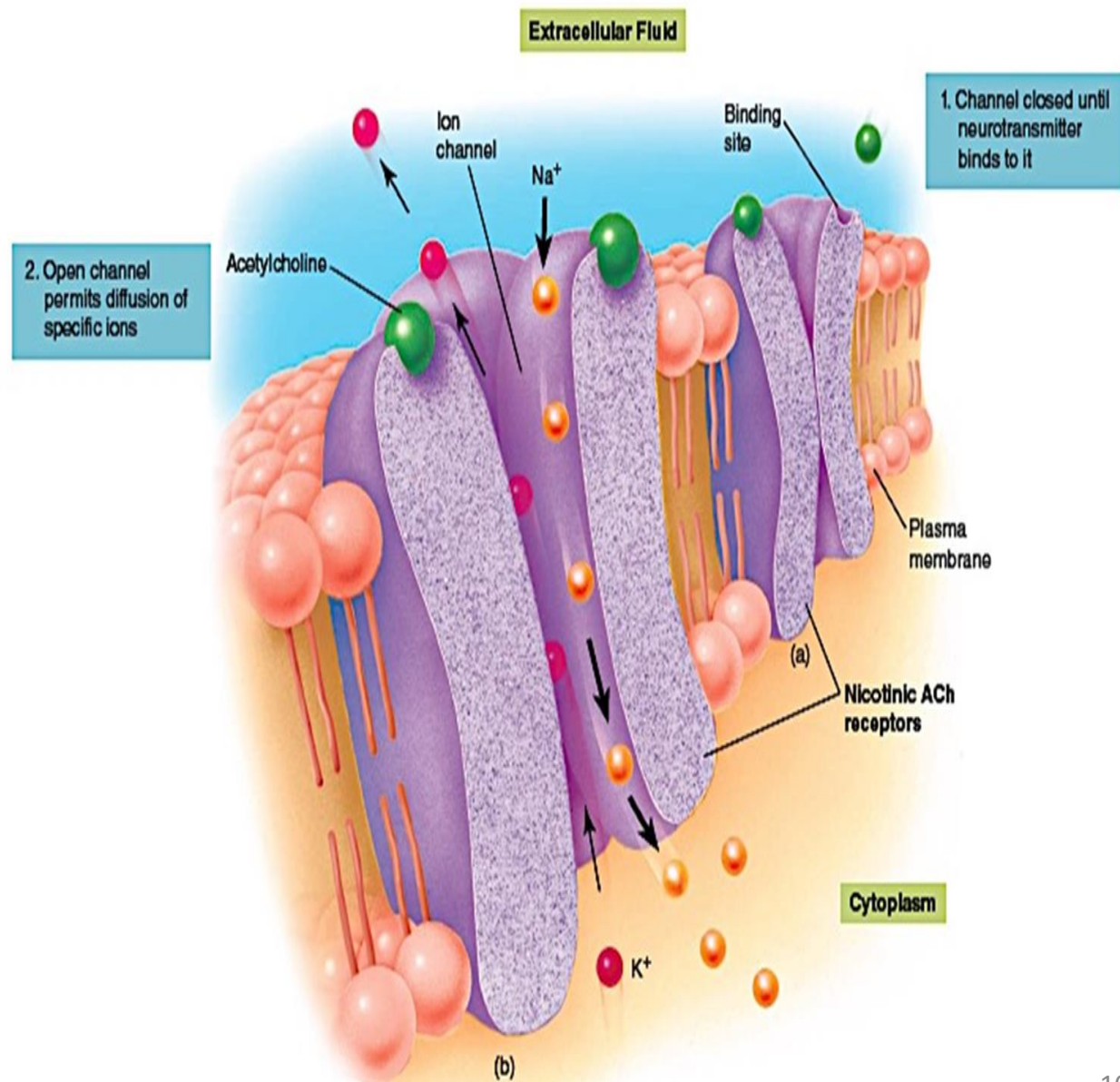
# Chemically Regulated Channels

- The majority of synapses in the nervous system is one-way and occurs through the release of chemical neurotransmitters from presynaptic axon endings.
- These presynaptic endings, called **terminal boutons**
- Neurotransmitters can open ion channels through binding to one of two types of receptors:
  - A. Ligand-gated ion channels receptors
  - B. G-protein coupled receptors



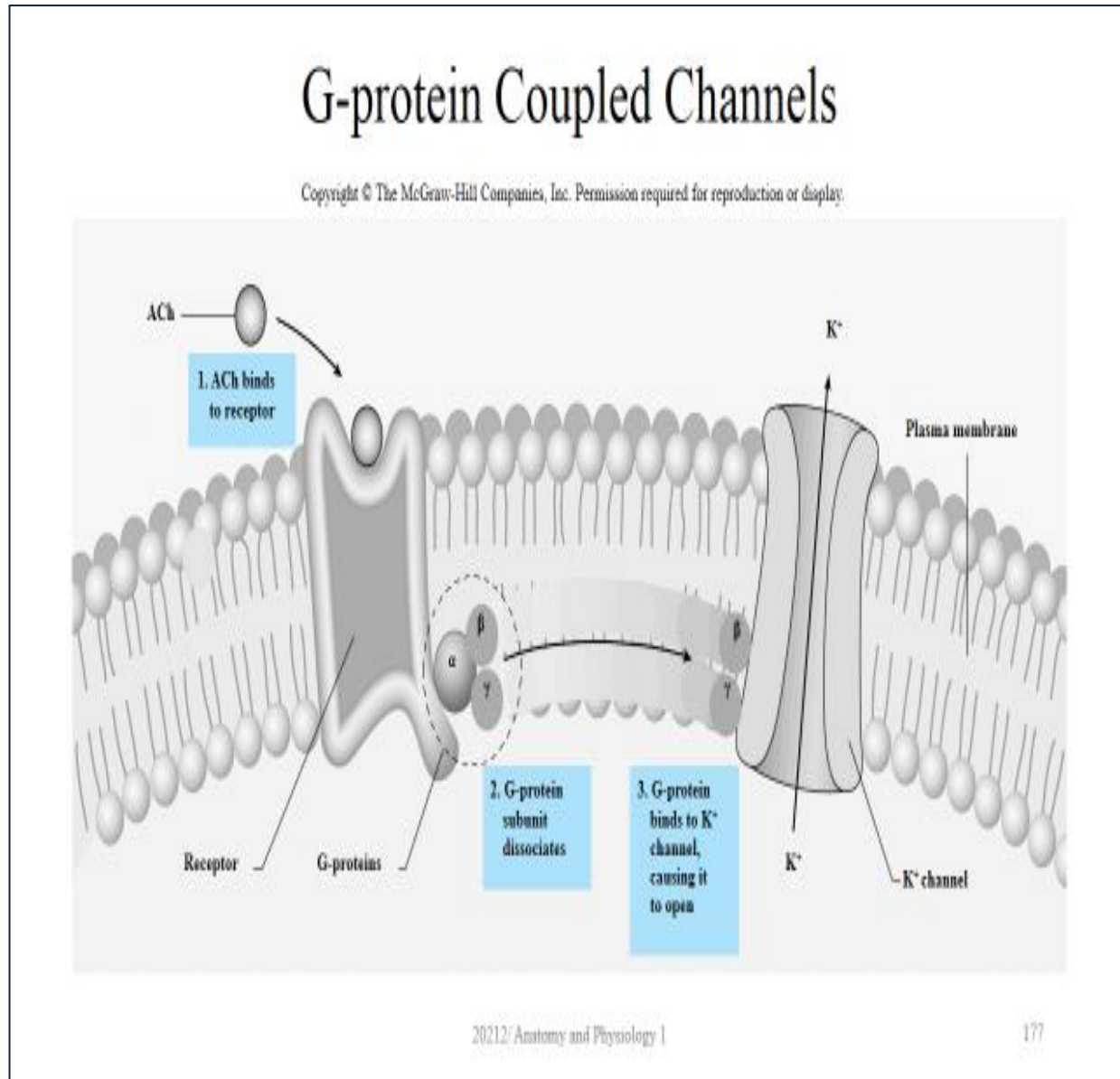
# Ligand-Gated Channels

- a. The **receptor protein** is also an **ion channel**; binding of the neurotransmitter directly opens the ion channel.
- b. **Nicotinic ACh receptors** are ligand-gated channels with two receptor sites for two AChs.
- c. Binding of 2 acetylcholine molecules opens a channel that allows both  $\text{Na}^+$  and  $\text{K}^+$  passage.
  - 1)  $\text{Na}^+$  flows in, and  $\text{K}^+$  flows out.
  - 2) Due to electrochemical gradient, more  $\text{Na}^+$  flows in than  $\text{K}^+$  out.
- d. This inward flow of  $\text{Na}^+$  depolarizes the cell, creating an EPSP.
  - 1) **EPSPs** occur in the dendrites and cell bodies.
  - 2) **EPSPs** from the binding of several ACh molecules can be added together to produce greater depolarization – graded depolarization
  - 3) **This** may reach the threshold for voltage-gated channels in the axon hillock, leading to action potential.



# G-Protein Coupled Channels

- a. The neurotransmitter receptor is separate from the protein that serves as the ion channel.
  - 1) Binding at the receptor opens ion channels indirectly by using a G-protein.
  - 2) **Muscarinic ACh receptors** interact with ion channels in this way as well as **dopamine** and **norepinephrine receptors**
- b. **Associated** with a G-protein
  - 1) G-proteins have three subunits (alpha, beta, and gamma).
  - 2) Binding of one acetylcholine results in the dissociation of the alpha subunit.
  - 3) Either the alpha or the beta-gamma diffuses through the membrane to the ion channel.
  - 4) This opens the channel for short period of time.
  - 5) The G-protein subunits dissociate from the channel, and it closes.



# G-Protein Coupled Channels

c. Binding of acetylcholine opens  $K^+$  channels in some tissues (IPSP) or closes  $K^+$  channels in others (EPSP).

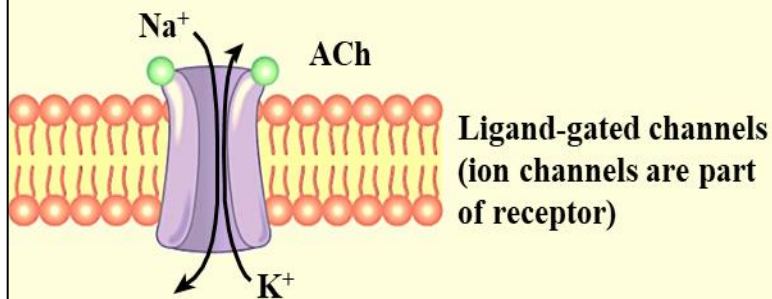
- 1) **In the heart**,  $K^+$  channels are opened by the beta-gamma complex, creating IPSPs (hyperpolarization) that slow the heart rate.
- 2) **In the smooth muscles of the stomach**,  $K^+$  channels are closed by the alpha subunit, producing EPSPs (depolarization) and the contraction of these muscles.

# Comparison of Nicotinic & Muscarinic ACh Receptors

## Nicotinic ACh receptors

Postsynaptic membrane of

- All autonomic ganglia
- All neuromuscular junctions
- Some CNS pathways

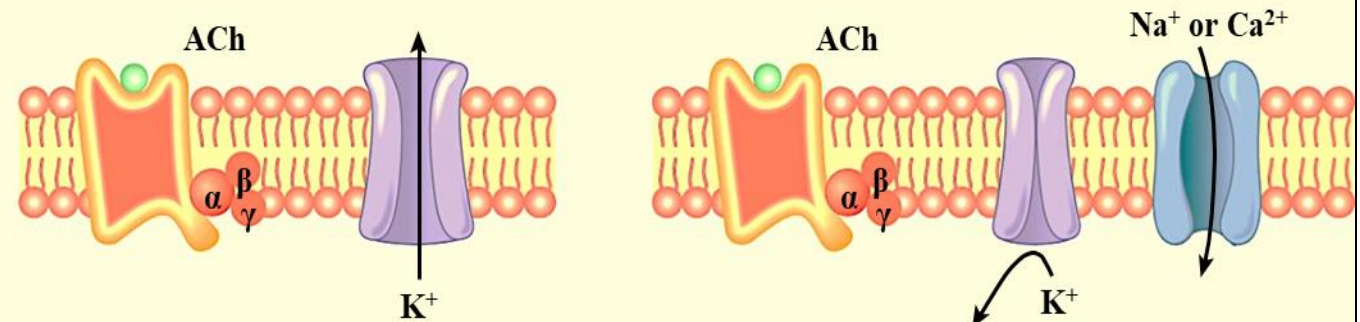


Depolarization

Excitation

## Muscarinic ACh receptors

- Produces parasympathetic nerve effects in the heart, smooth muscles, and glands
- G-protein-coupled receptors (receptors influence ion channels by means of G-proteins)



Hyperpolarization

(K<sup>+</sup> channels opened)

Inhibition

Produces slower heart rate

Depolarization

(K<sup>+</sup> channels closed)

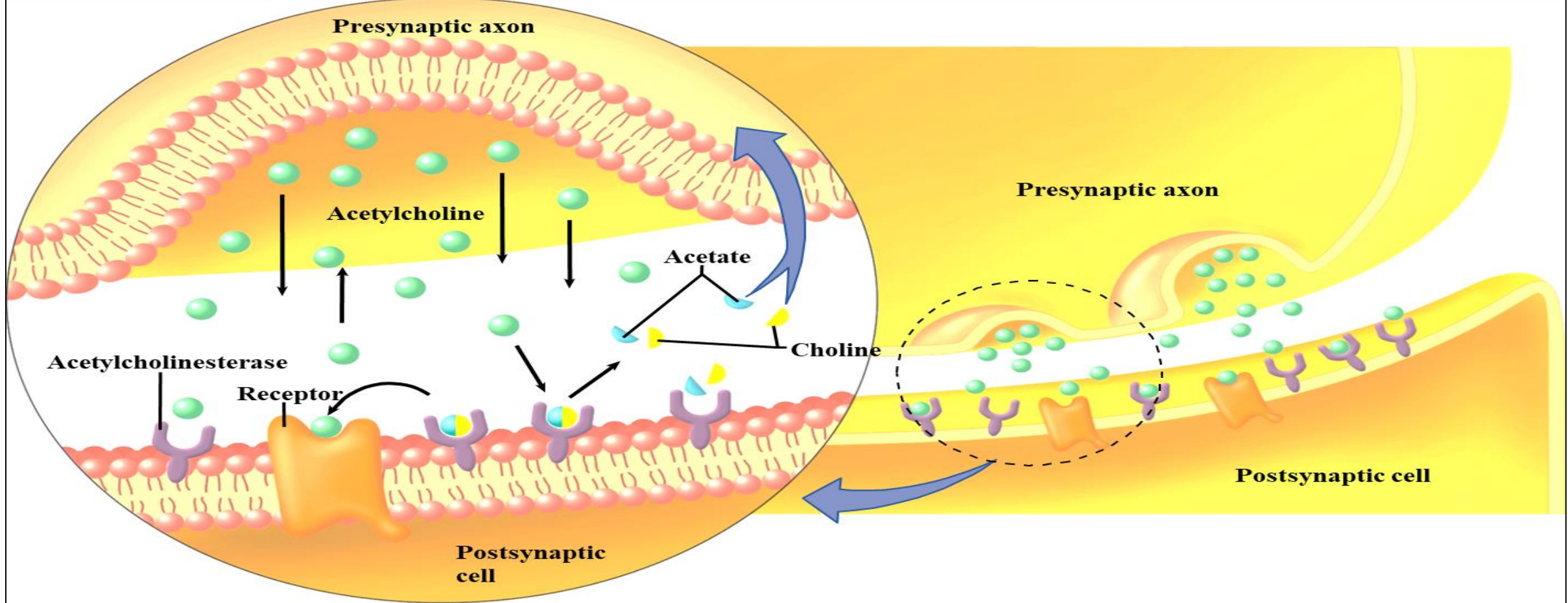
Excitation

Causes smooth muscles of the digestive tract to contract



# Action of Acetylcholinesterase (AChE)

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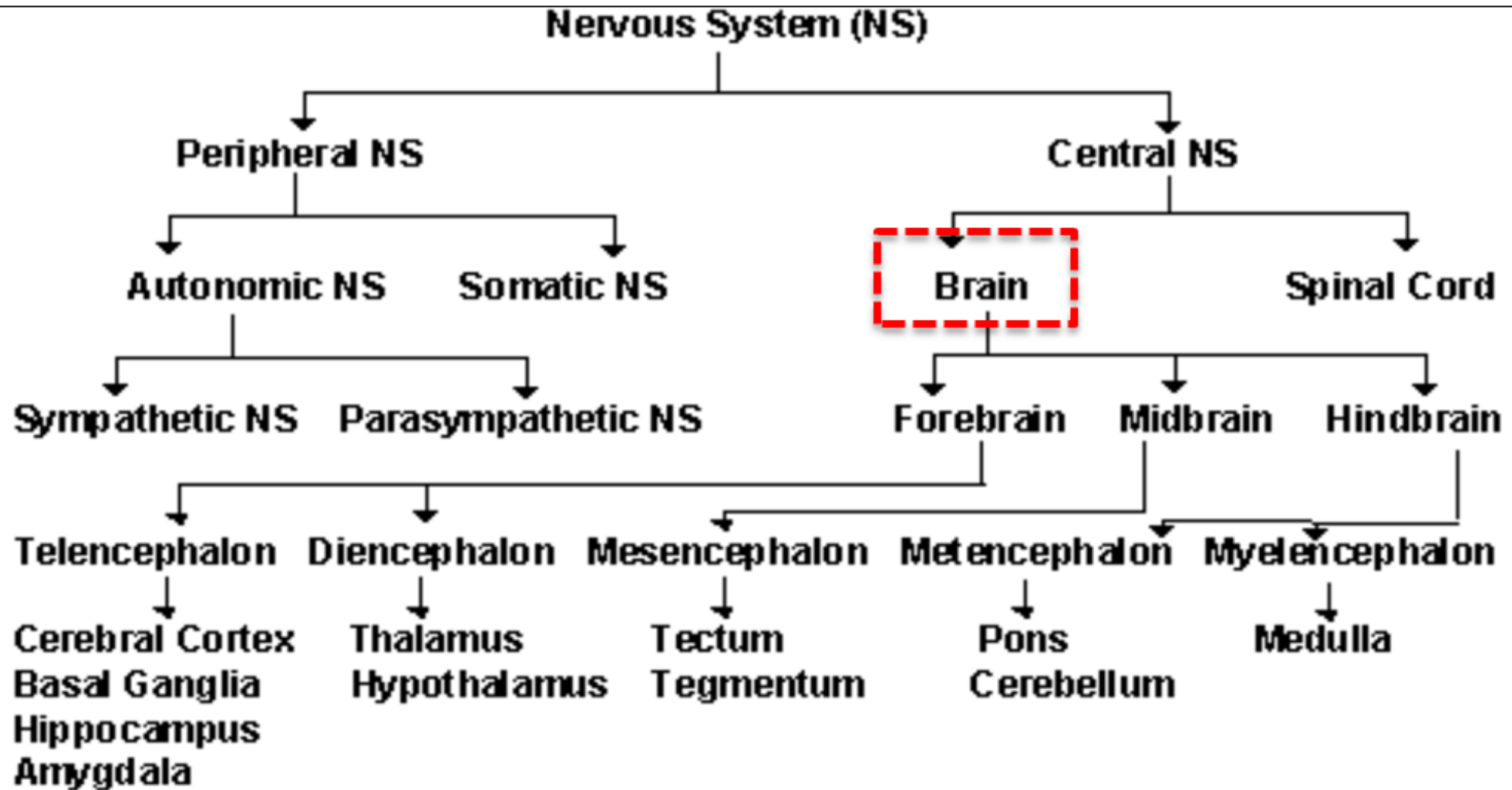
- To stop the activity in the postsynaptic cell
- **AChE** is an enzyme that inactivates ACh activity shortly after it binds to the receptor.
- Hydrolyzes ACh into acetate and choline, which are taken back into the presynaptic cell for reuse (reuptake).

# **Chapter 8:**

## **Central nervous system (CNS)**



## Organization of The NS

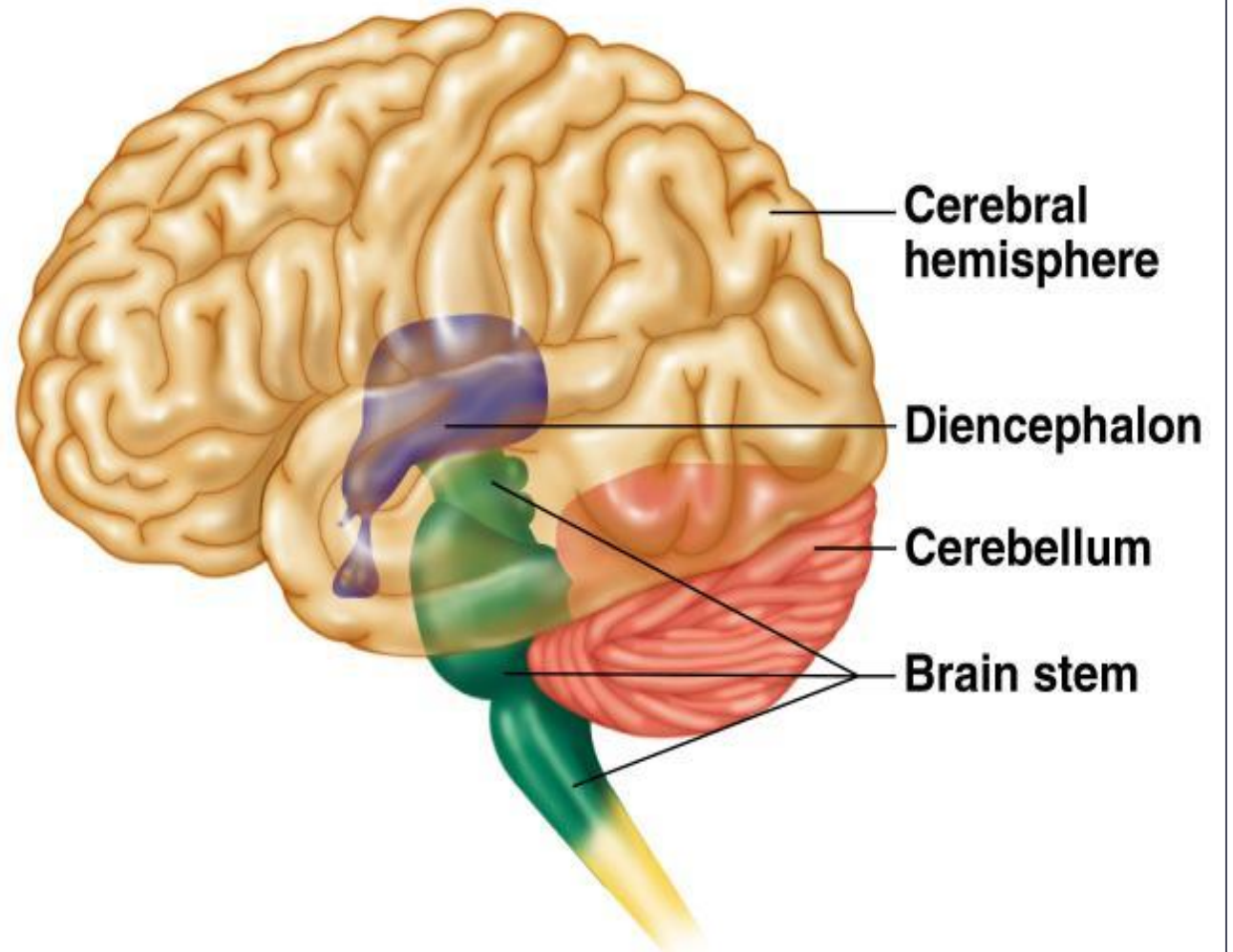


## Chapter 8: Central Nervous System

- ✧ The central nervous system (CNS), consisting of the brain and spinal cord.
- ✧ It receives input from sensory neurons and directs the activity of motor neurons that innervate muscles and glands.
- ✧ The association neurons within the brain and spinal cord are in a position, associate appropriate motor responses with sensory stimuli, and thus to maintain homeostasis in the internal environment and the continued existence of the organism in a changing external environment.
- ✧ Perceptions, learning, memory, emotions, and perhaps even the self-awareness that forms the basis of consciousness, are creations of the brain

# Regions of the Brain

1. Cerebrum (cerebral hemispheres)
2. Diencephalon
3. Brain stem
4. Cerebellum

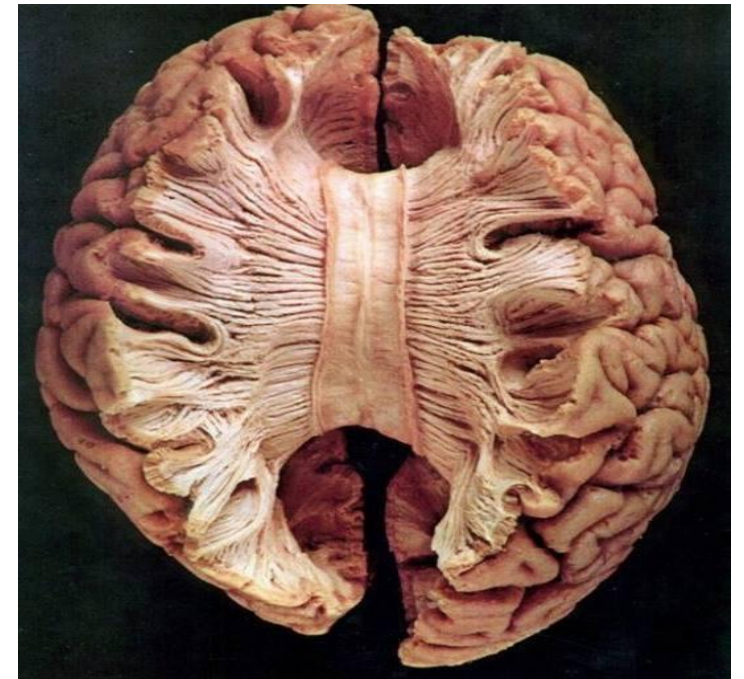


# The **Cerebrum**: Introduction

1. **The Cerebrum** is derived from the **telencephalon** and is the only structure of the telencephalon.
2. **Largest portion** of the brain (80% of the mass).
3. **Responsible for higher** mental functions.
4. **Consists of a right and left cerebral hemisphere** connected internally by a large fiber tract called the **corpus callosum**.



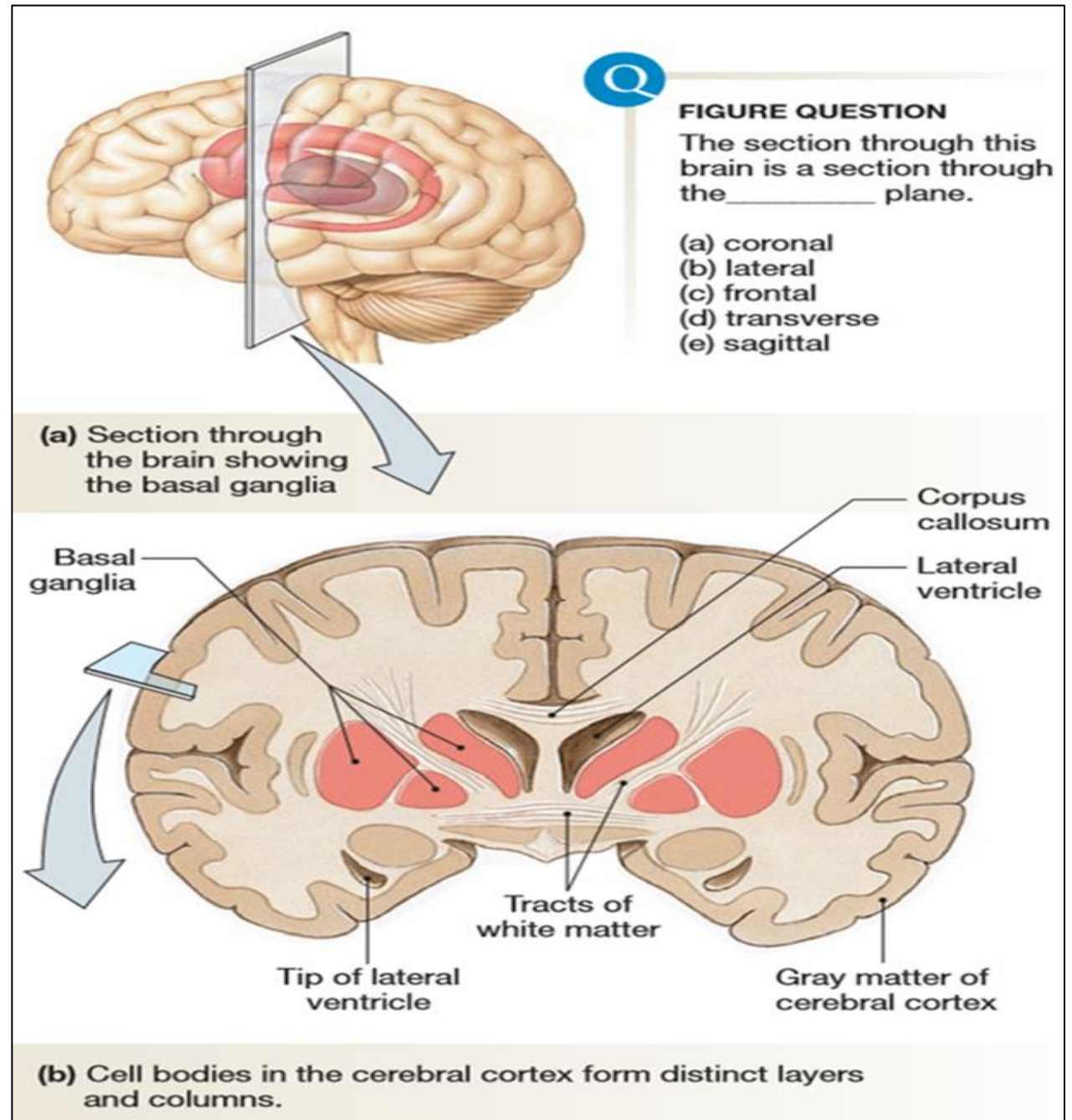
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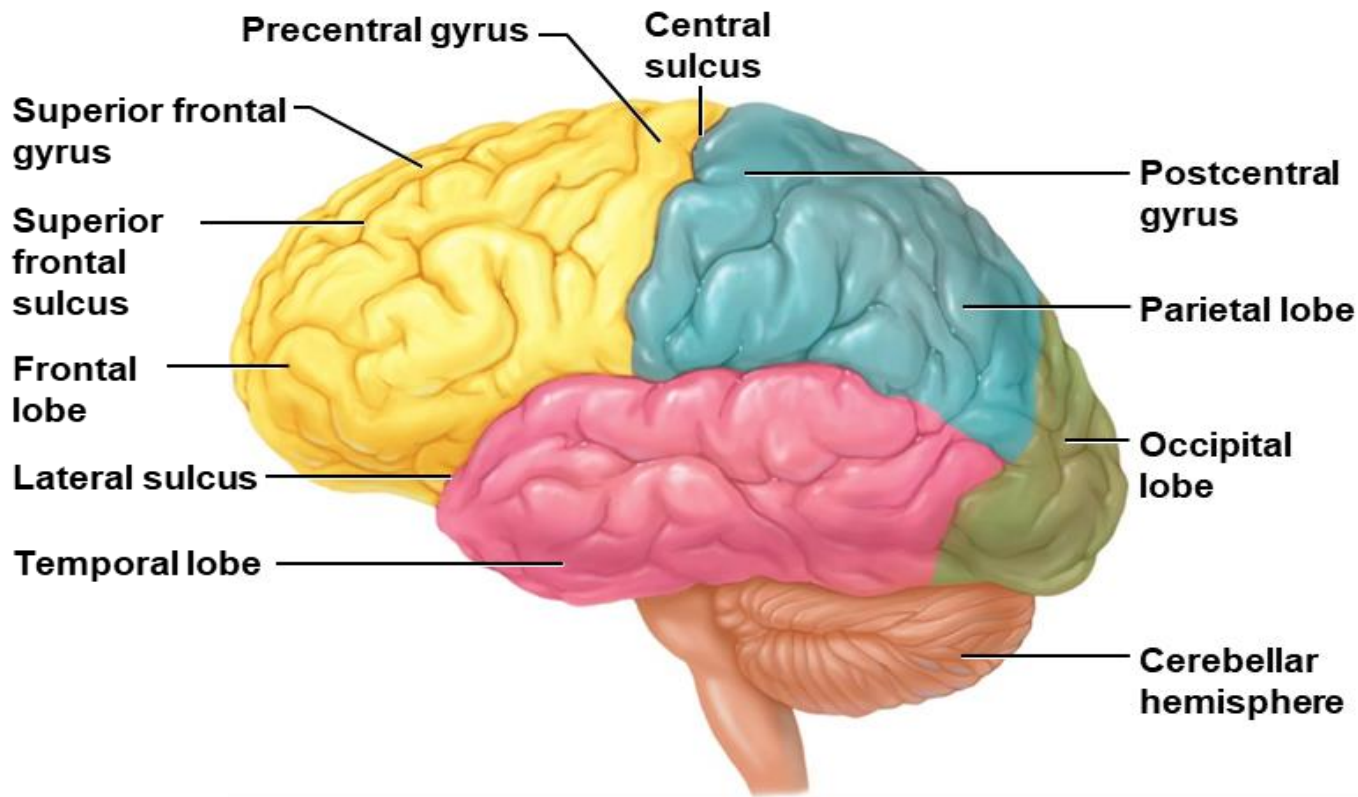
# Cerebral Cortex

1. The outer region of the cerebrum is called the **cerebral cortex** and is composed of 2–4 mm **gray matter** with underlying **white matter**.
2. Characterized by raised folds called **gyri** separated by depressed grooves called **sulci**; together called convolutions

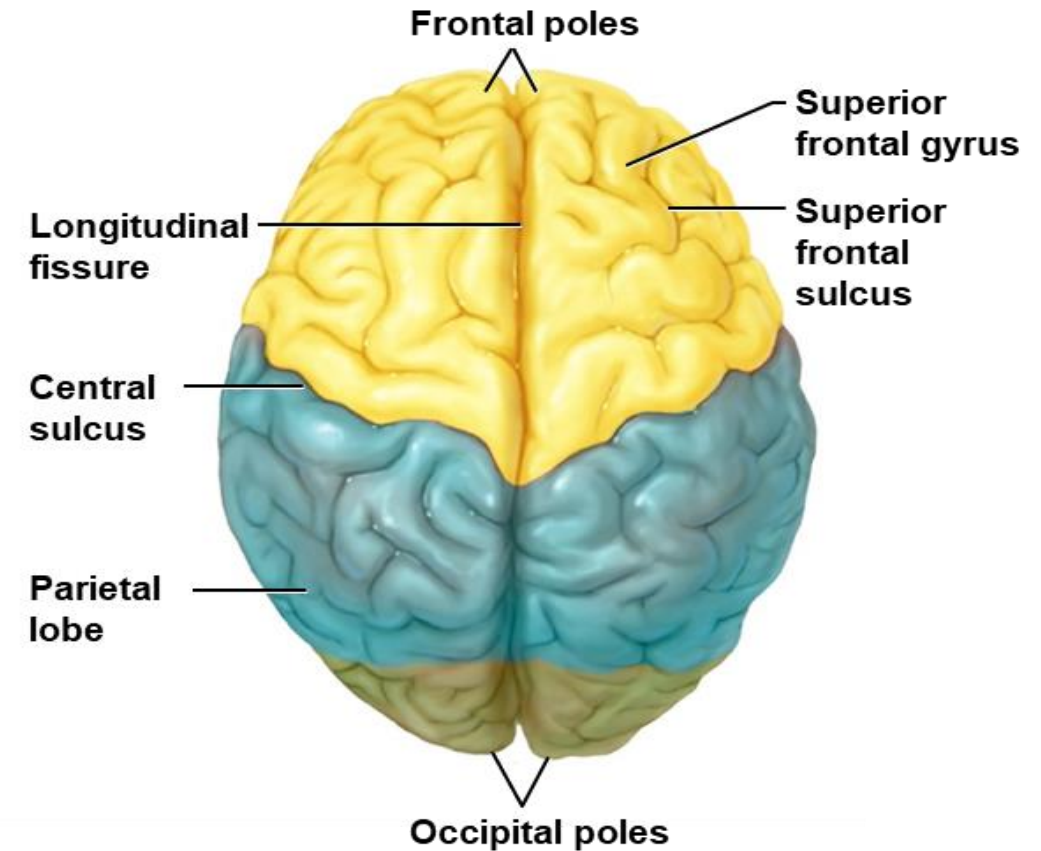


# Lobes of the Cerebrum

- Each hemisphere is divided by **deep sulci** or **fissures** into 5 lobes - **Frontal** , **Parietal**, **Temporal**, **Occipital**, **Insula**



(a)

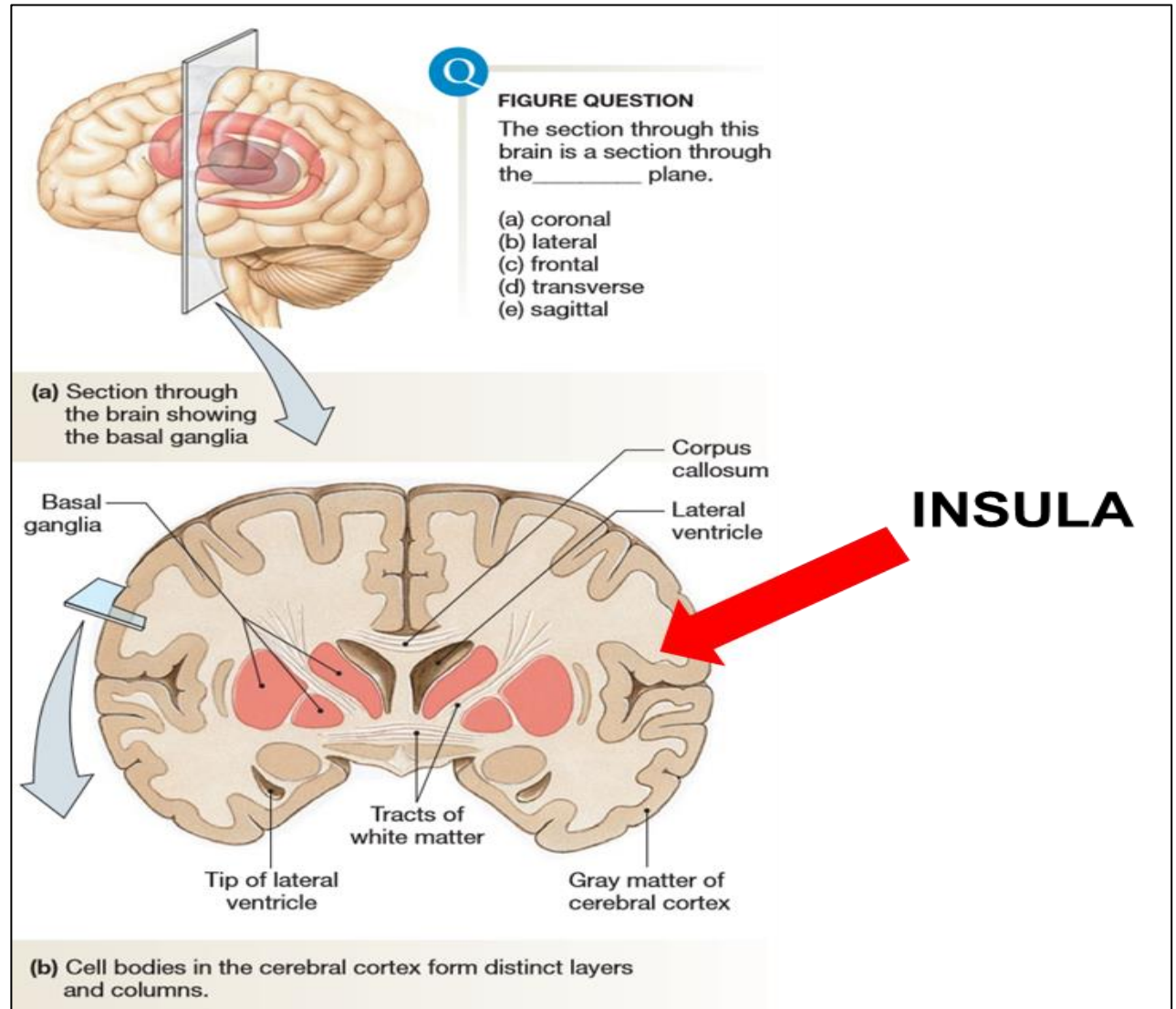
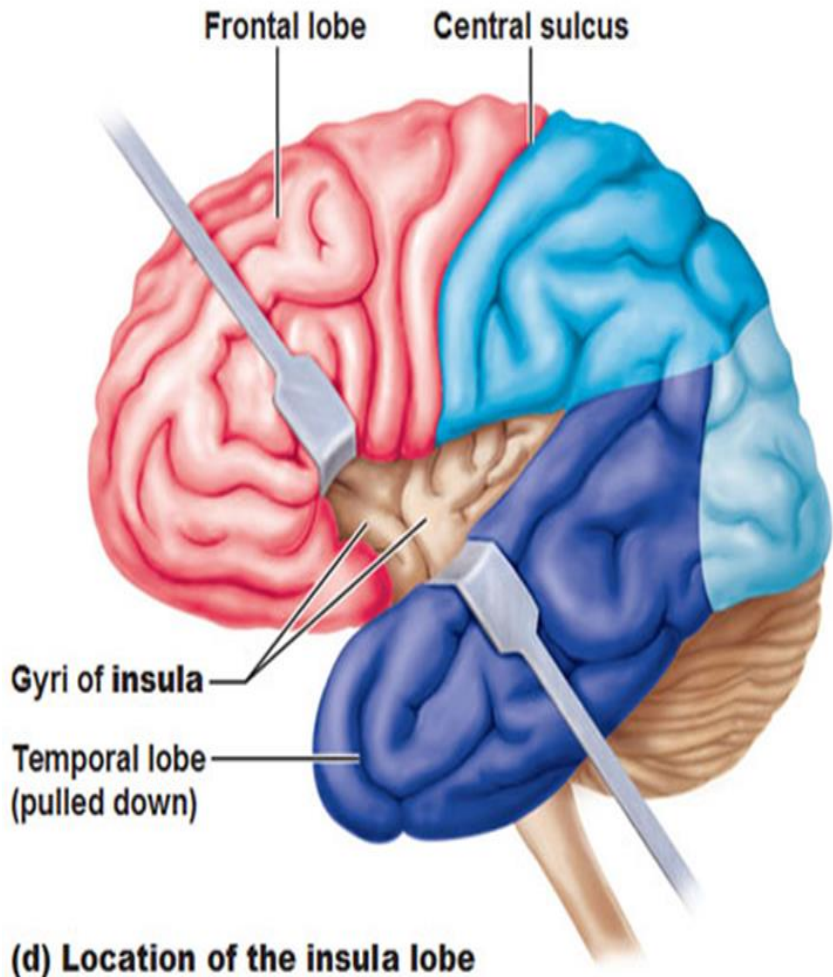


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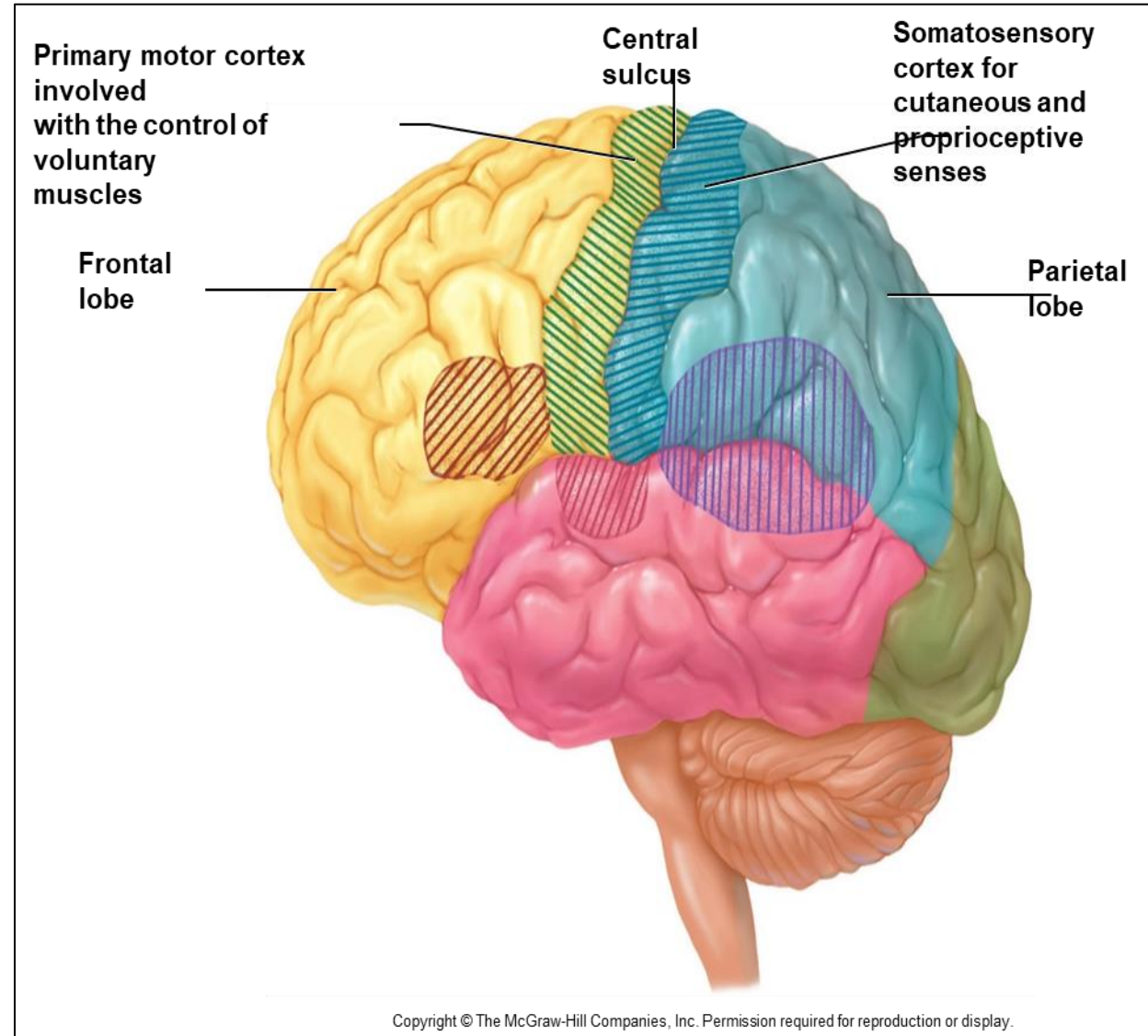
# Lobes of the Cerebrum: the **Insula**

## The Cerebral Hemispheres – one more lobe



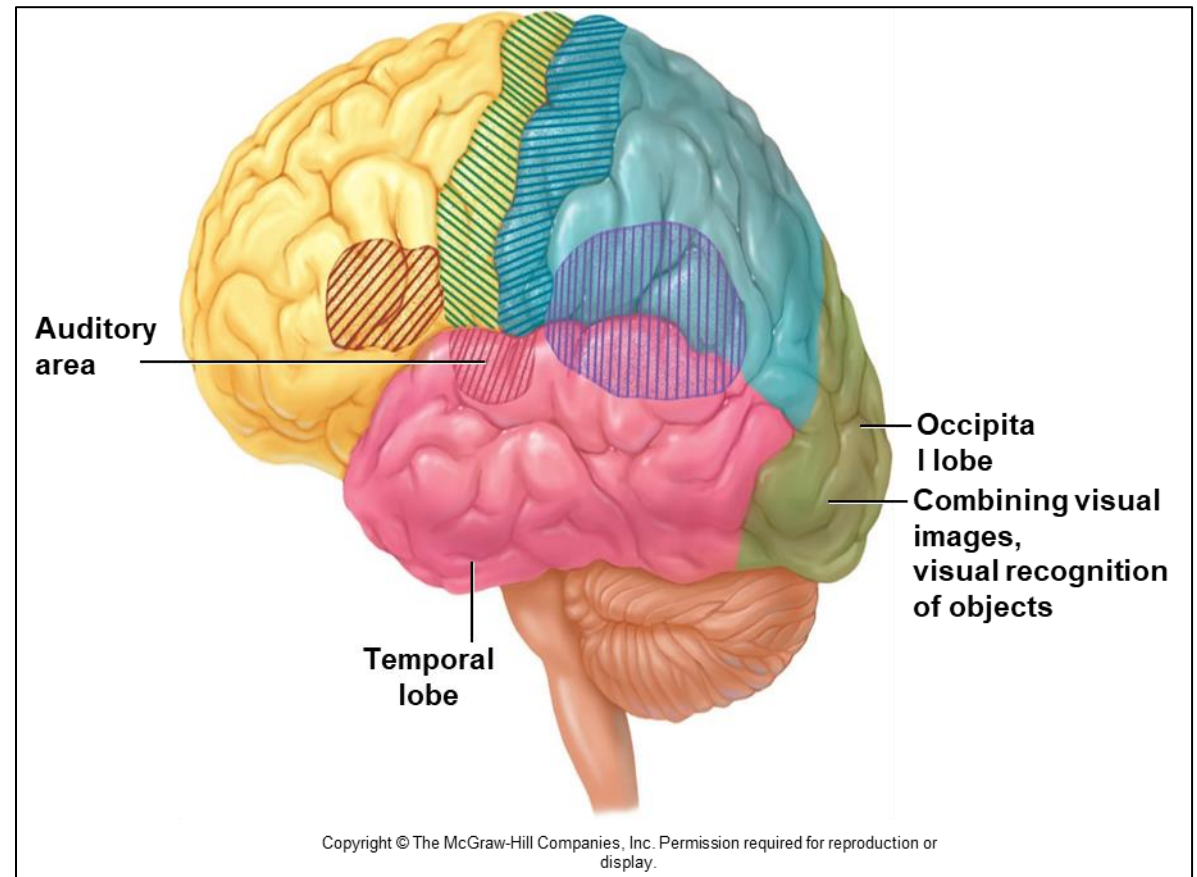
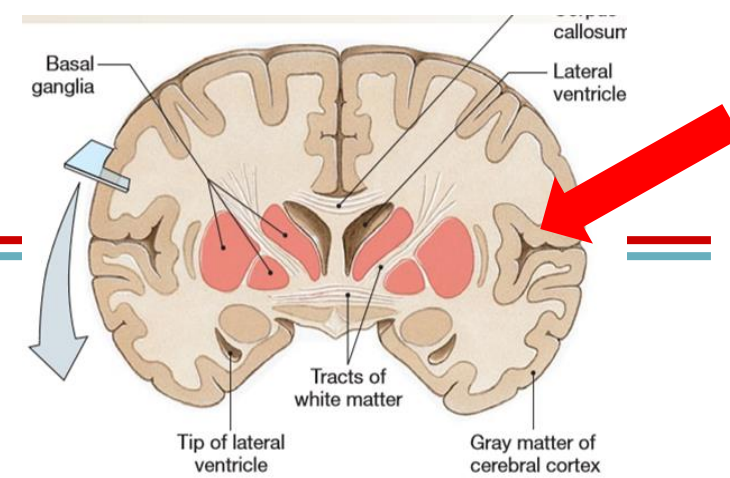
# Frontal and Parietal Lobes

- The frontal lobe is located at the front of the brain and the parietal lobe is located in the middle section of the brain
- Separated by the **central sulcus**
- The **precentral gyrus** is located in the **frontal lobe** and is responsible for motor control (also called **primary motor cortex**) ; the cell bodies of the interneurons in this area are called upper motor neurons because of their role in muscle regulation.
- The **postcentral gyrus** is in the **parietal lobe** and is responsible for somesthetic sensation (coming from receptors in the skin, muscles, tendons, and joints); called the **somatosensory cortex**



# Temporal, Occipital, and Insula Lobes

- The temporal lobe is located on the **bottom section of the brain** and the occipital lobe is located at the **back portion of the brain**.
- **Temporal lobe:** auditory centers (receives sensory fibers from the cochlea).
- **Occipital lobe:** responsible for vision and coordination of eye movements
- **Insula:** encoding of memory and integration of sensory information with visceral responses; receives olfactory, gustatory (taste), auditory, and pain information; important in assessing the bodily states that accompany emotions.



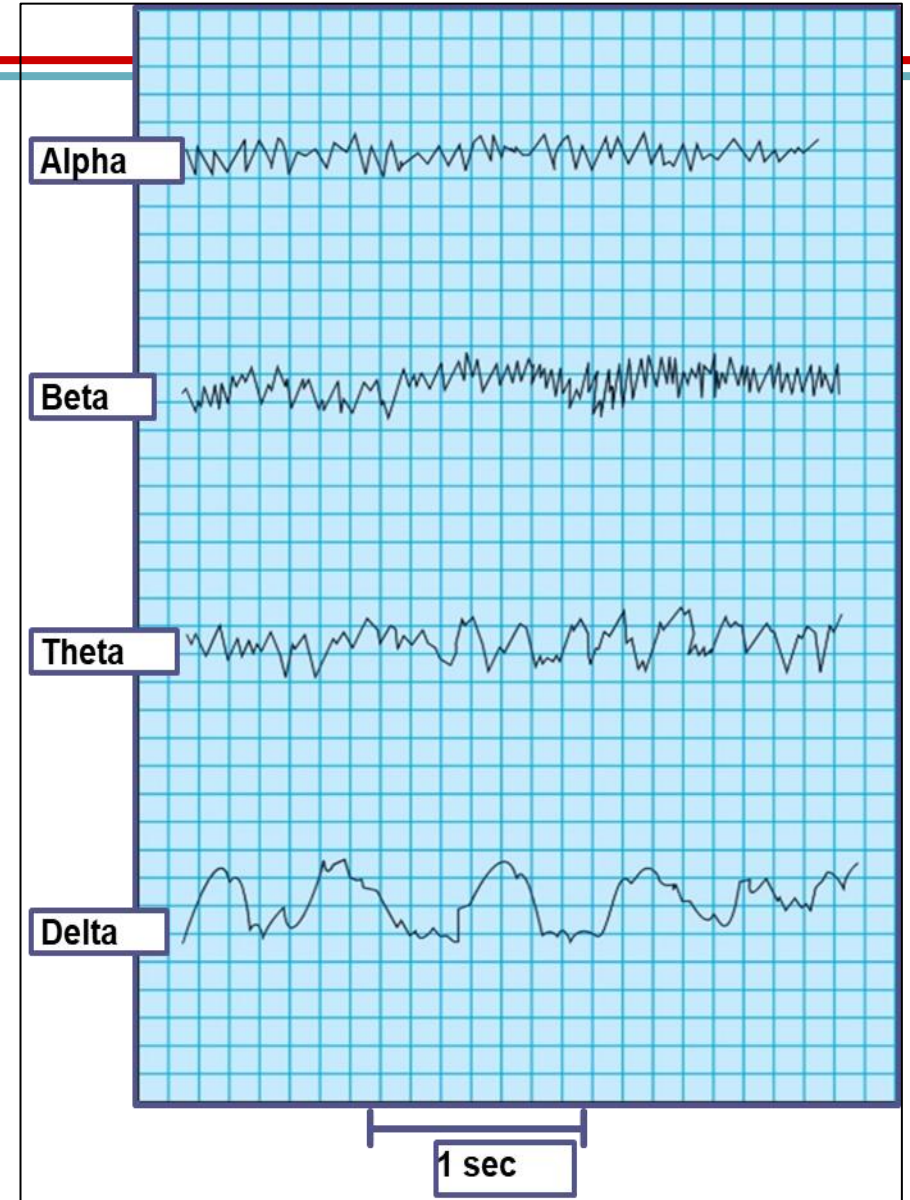


# Visualizing the Brain

**Electroencephalogram (EEG):** Electrodes on the scalp detect synaptic potentials produced by cell bodies and dendrites in the cerebral cortex.

**Four patterns are usually seen:**

- **Alpha waves:** active, relaxed brain. Seen most in frontal and parietal lobes.
- **Beta waves:** produced with visual stimulation and mental activity. Seen most in frontal lobe.
- **Theta waves:** seen during sleep; most from occipital and temporal lobes.
- **Delta waves:** also seen in sleep, from all over the cerebrum.



# Sleep

- May be genetically controlled, although sleep is affected by environmental factors
- Two recognized categories:
  - 1) **REM**: rapid eye movement; state when dreams occur. Theta waves are seen here.
  - 2) **Non-REM**: also called resting sleep; divided into four stages, determined by EEG waves seen. Stages 3 and 4 are often called slow-wave sleep, characterized by delta waves.

# Sleep pattern

- 1) **When people first fall asleep**, they enter non-REM sleep and progress through the four stages.
- 2) **Next**, a person ascends back up the stages of non-REM sleep to REM sleep.
- 3) **This cycle repeats** every 90 minutes, and most people go through five per night.
- 4) **If allowed to awaken naturally**, people usually do so during REM sleep.
- 5) **Slow-wave is prominent** in the first part of sleep, while REM is prominent in the second half



## REM Sleep

- 1) **Some brain regions** are more active during REM sleep than during the waking state.
- 2) **The limbic system** (involved in emotion) is very active during REM sleep.
- 3) **Breathing and heart** rate may be very irregular.
- 4) **Benefits consolidation** of nondeclarative memories (long term non-conscious memory).

## Non-REM Sleep

- 1) As you fall asleep, neurons decrease their firing rates, decreasing blood flow and energy metabolism.
- 2) Breathing and heart rate are very regular.
- 3) Non-REM sleep may allow repair of metabolic damage done to cells by free radicals and allows time for the neuroplasticity mechanisms needed to store memories.
- 4) Benefits consolidation of spatial and declarative memories (long term conscious memory).

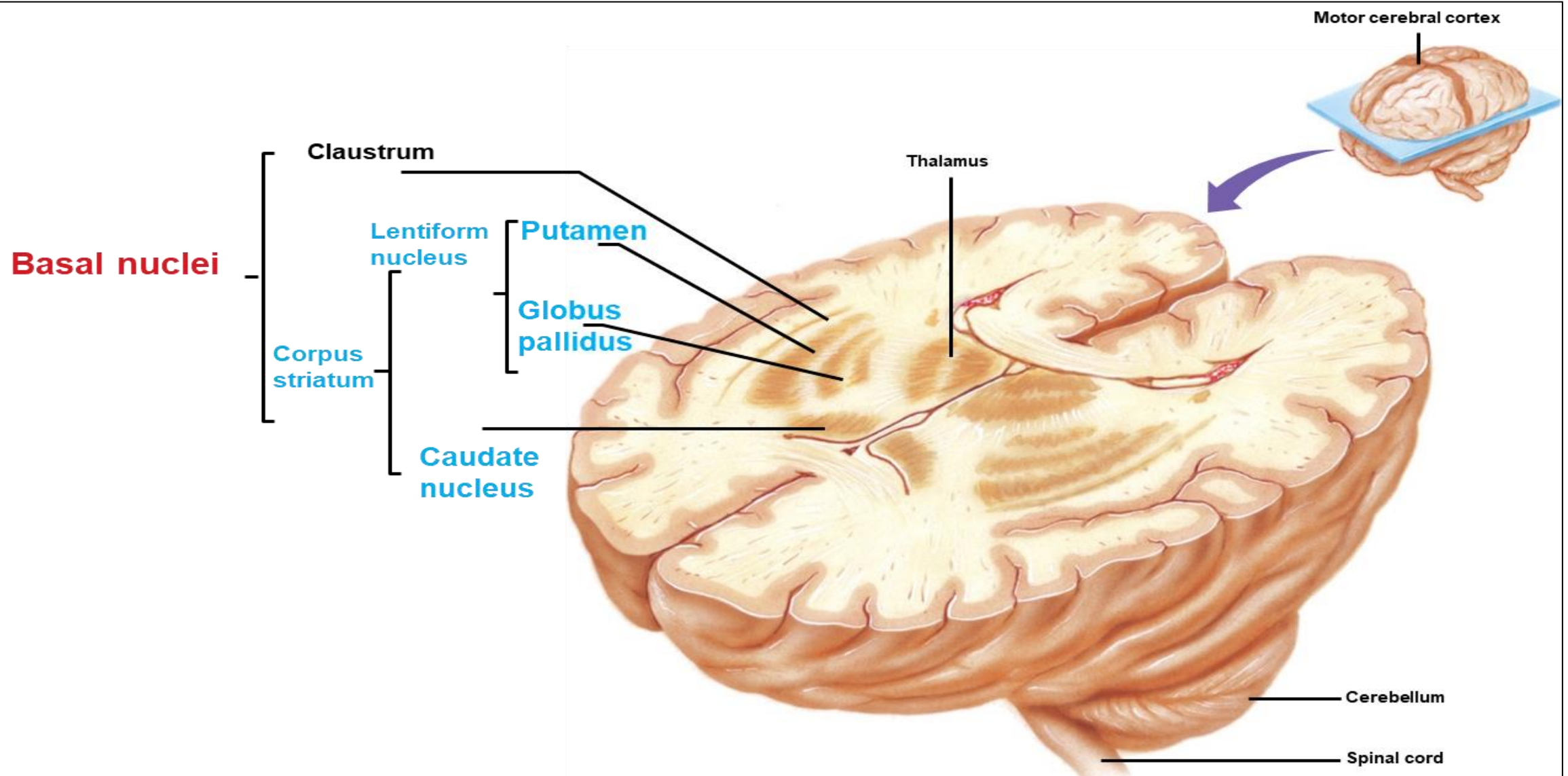
# Types of Memory

- a. **Short-term memory:** recent events; transferred to long-term memory through process of memory consolidation
  - 1) Memory consolidation occurs in the medial temporal lobe, hippocampus, and amygdala.
  - 2) Sleep is needed for optimum memory consolidation.
- b. **Long-term memory**
  - 1) Requires actual structural change - Activation of genes, synthesis of mRNA, production of proteins, and formation of new synapses
  - 2) Long-term memory can be classified into:
    - a) Nondeclarative (implicit): memory of simple skills, how to do things
    - b) Declarative (explicit): memory of things that can be verbalized. People with amnesia have impaired declarative memory; further broken into:
      - 1) **Semantic:** facts
      - 2) **Episodic:** events

## Basal Nuclei

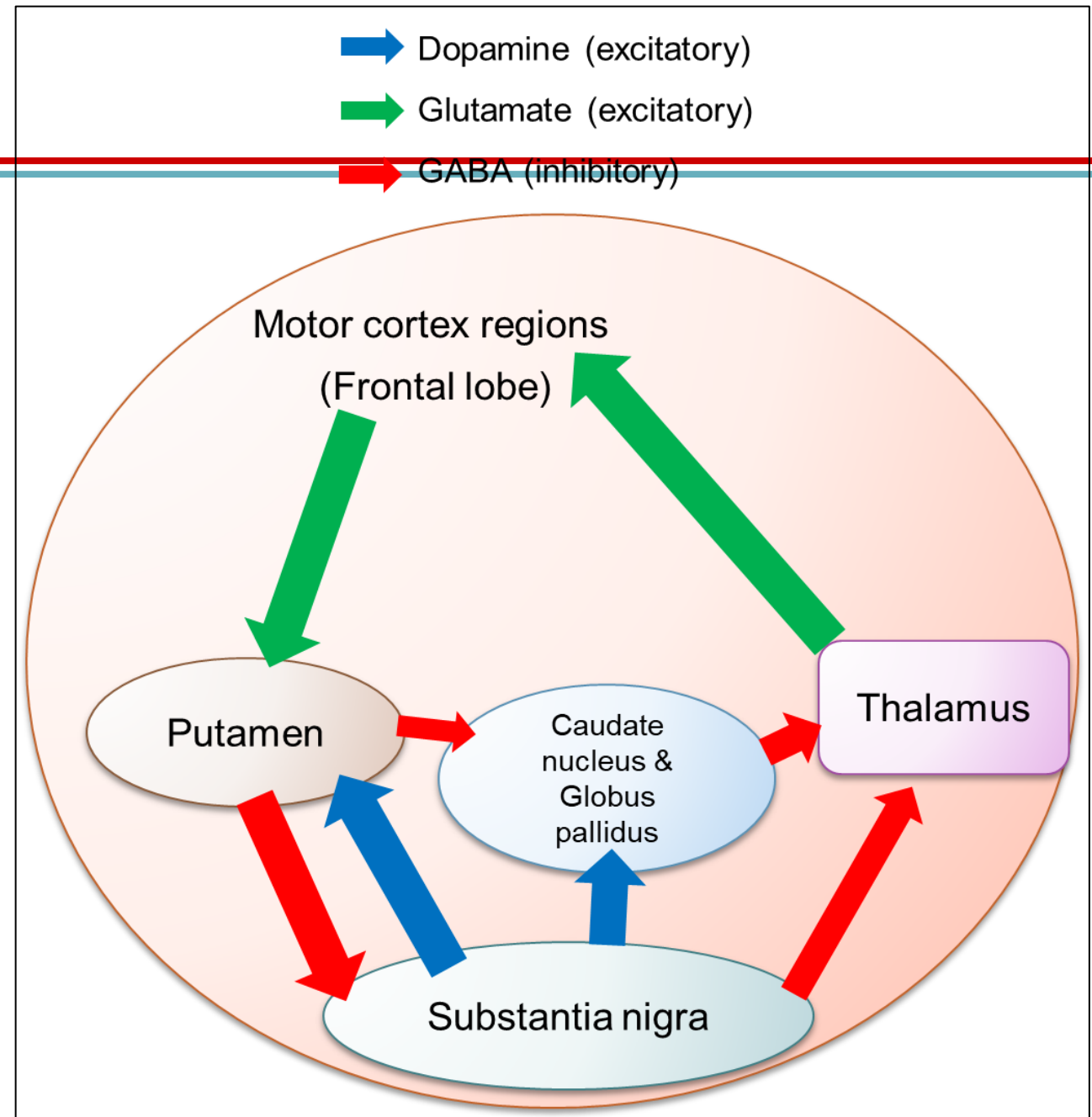
1. **Masses** of gray matter (neuron cell bodies) located deep in the white matter of the cerebrum
2. **Most** prominent is the **corpus striatum**; composed of:
  - a. **Caudate nucleus**
  - b. **Lentiform nucleus**; made up of the **putamen** and the **globus pallidus**.
3. Also includes **subthalamic nucleus** of the diencephalon and **substantia nigra** of the midbrain.
4. The basal nuclei function in the **control of voluntary movements**.

# Basal Nuclei, cont



# Motor circuit

- a. **The neurons from motor** cortex regions of the frontal lobe release glutamate (stimulatory) in the putamen. The putamen then releases GABA (inhibitory) to other regions of the basal nuclei.
- b. **The globus pallidus** and the substantia nigra sends GABA-releasing (inhibitory) neurons to the thalamus, which sends excitatory axons to the motor cortex of the cerebrum.
- c. This completes a **motor circuit**. This circuit **stimulates appropriate movements and inhibits unwanted movement**.
- **Substantia nigra**: Degeneration of **dopaminergic neurons** that project from the **substantia nigra** to the **corpus striatum (the nigrostriatal tract)** causes **Parkinson's disease**.



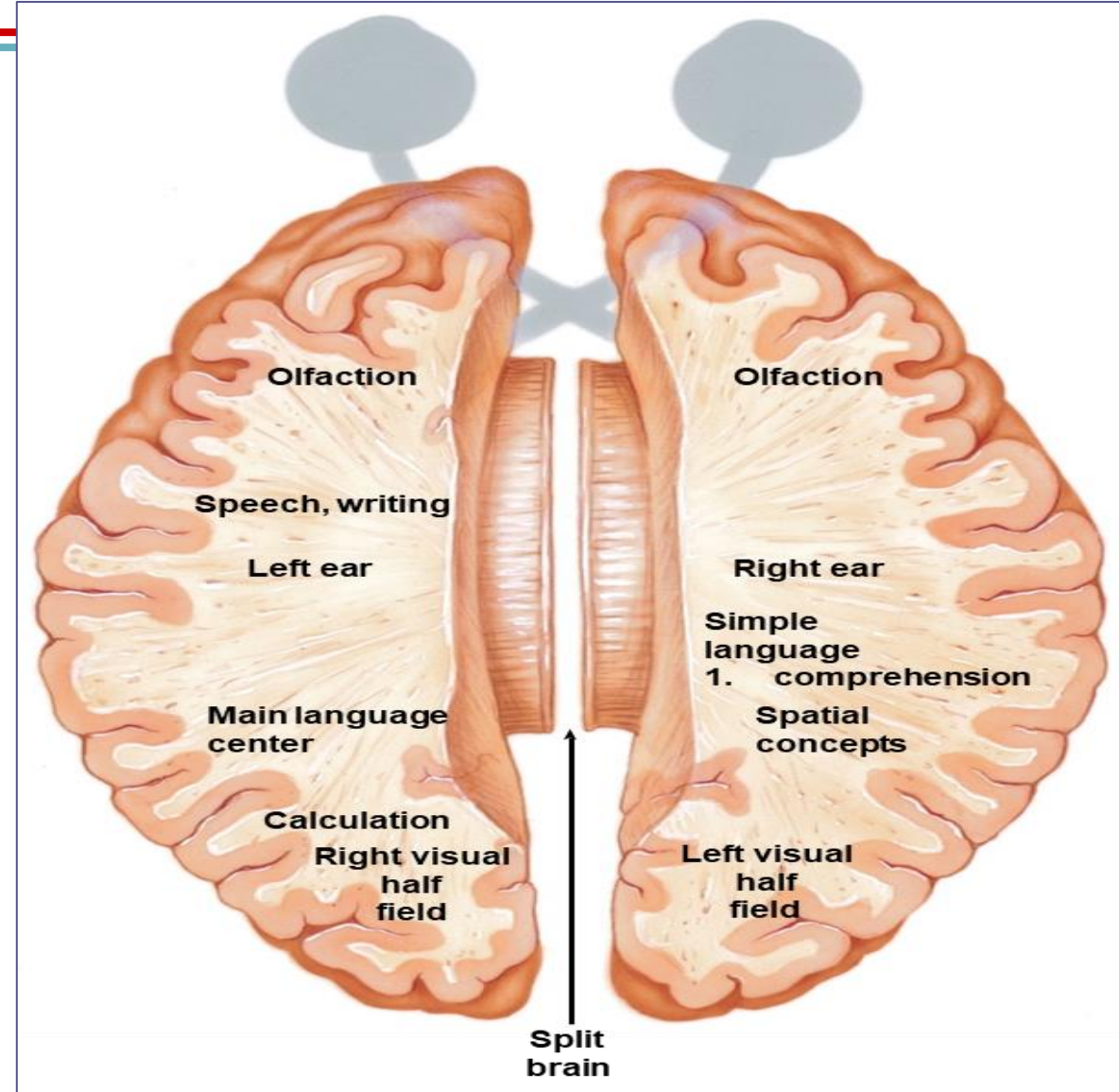


# Cerebral Lateralization vs. Cerebral Dominance

- Each side of the precentral gyrus controls movements on the contralateral (opposite) side of the body due to **decussation (crossing over) of fibers**.
- Somesthetic sensation from each side of the body projects to contralateral sides of the postcentral gyrus.
- Communication between the sides occurs through the **corpus callosum**; this is severed in severe forms of epilepsy.
- **Cerebral dominance** (analogous to **handedness**) suggests that people generally have greater competence with **one hemisphere** than with the other.
- **Cerebral lateralization** suggests that each hemisphere is **specialized** in different **functions**. Thus, the **two hemispheres** appear to have **complementary functions** rather than one hemisphere being dominant and the other subordinate.
- The term **cerebral lateralization**, or specialization of function in one hemisphere or the other, is thus now **preferred** to the term **cerebral dominance**, although **both terms are currently used**.

# Cerebral Lateralization

- **Cerebral lateralization** suggests that each hemisphere is **specialized** in different **functions**.
- Some tasks seem to be performed better by one side of the brain than the other.
  - a. **Right hemisphere:** visuospatial tasks, recognizing faces, arranging blocks, reading maps
  - b. **Left hemisphere:** Language, speech, writing, calculations, analytical abilities

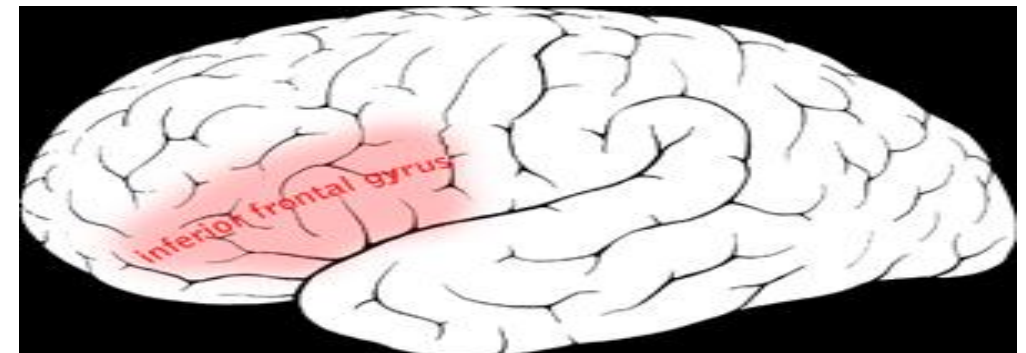
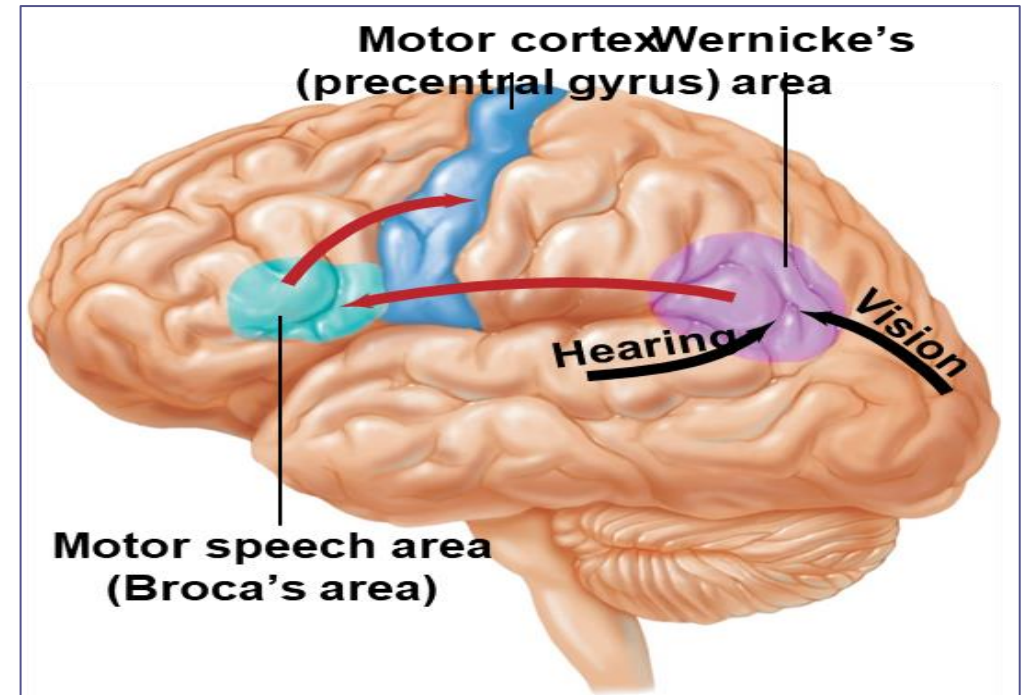


# Language

- Most of the knowledge of how the brain controls language has come from studying people with speech problems called **aphasias**.
- Aphasias are speech and language disorders caused by damage to the brain through head injury or stroke.
- Two areas are identified as important (the language areas of the brain are located in most people in the left hemisphere of the cerebral cortex):
  - a. **Broca's area**
  - b. **Wernicke's area**

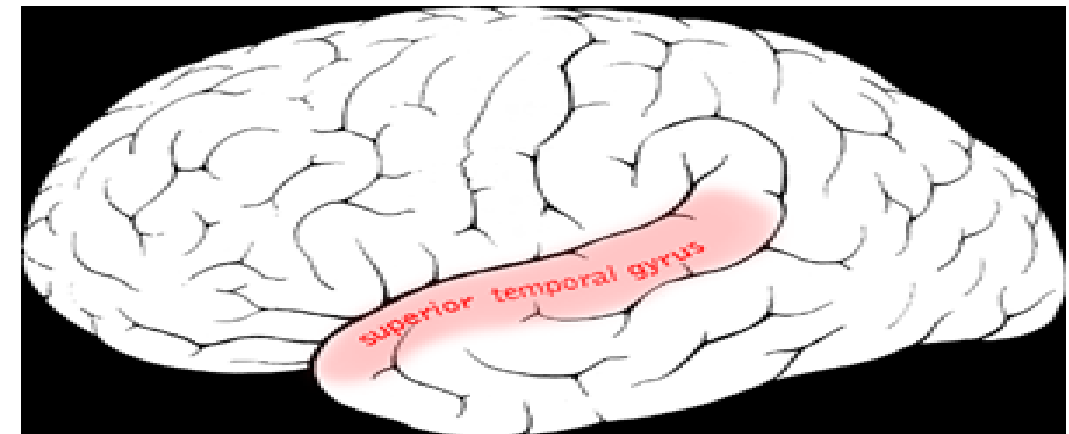
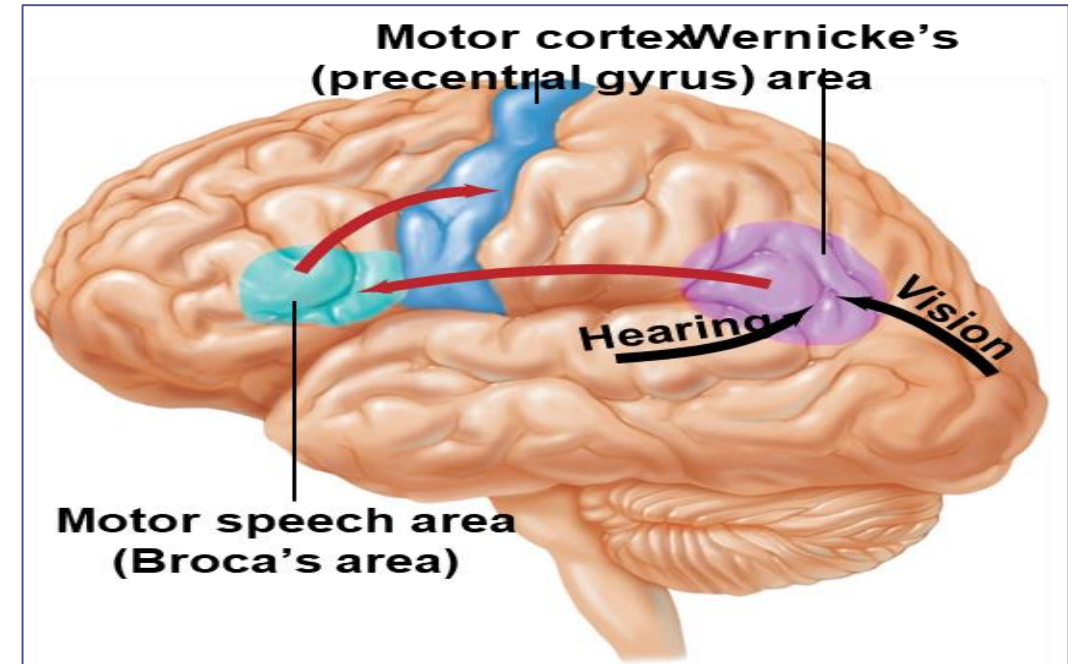
# Broca's Area

- a. **Located in left** inferior frontal gyrus
- b. **sends fibers to** the motor cortex (precentral gyrus), which directly controls the musculature of speech (Controls motor aspects of speech).



# Wernicke's Area

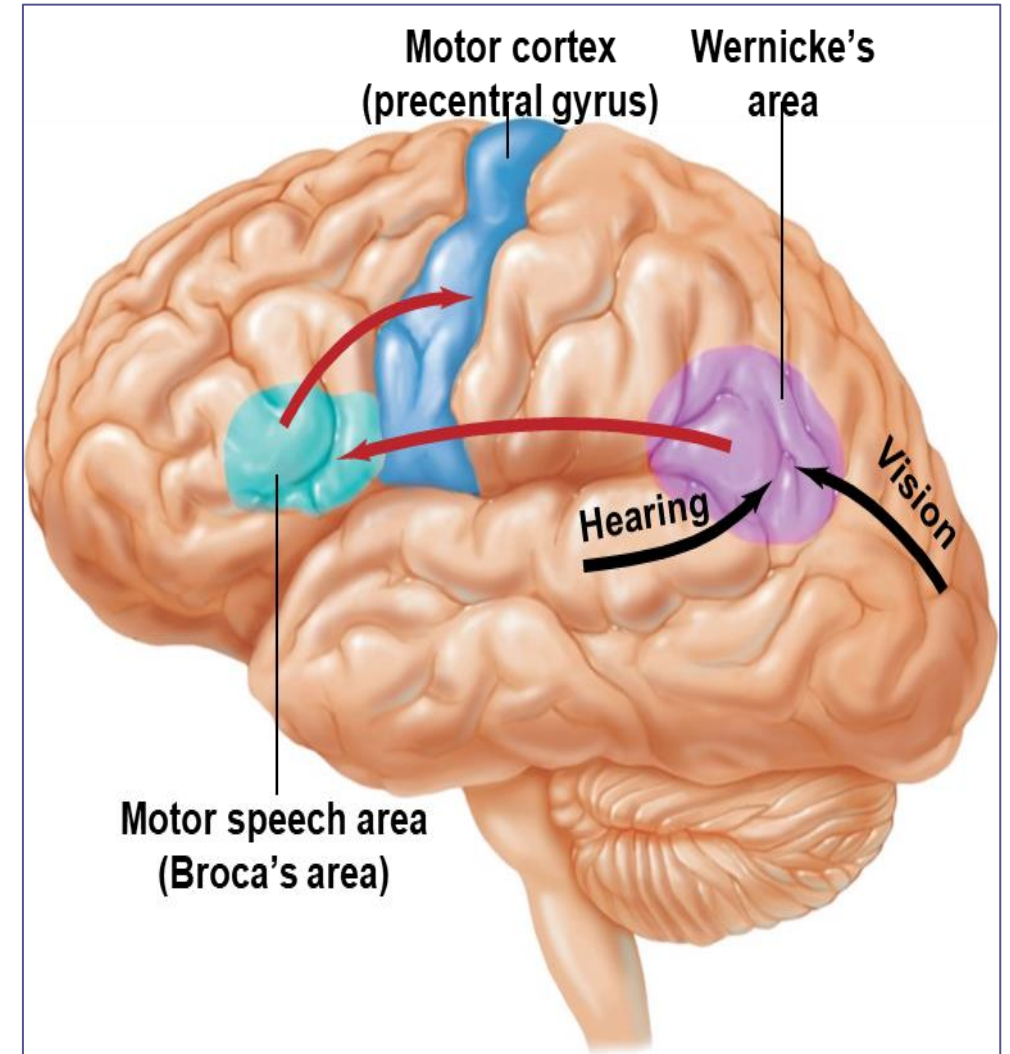
- a. **Located** in left superior temporal gyrus
- b. **Controls** understanding of words (language comprehension).
- c. **Information** about heard words is sent by the temporal lobe (auditory cortex).
- d. **Information** about written words is sent by the occipital lobe (visual cortex).





# Speech

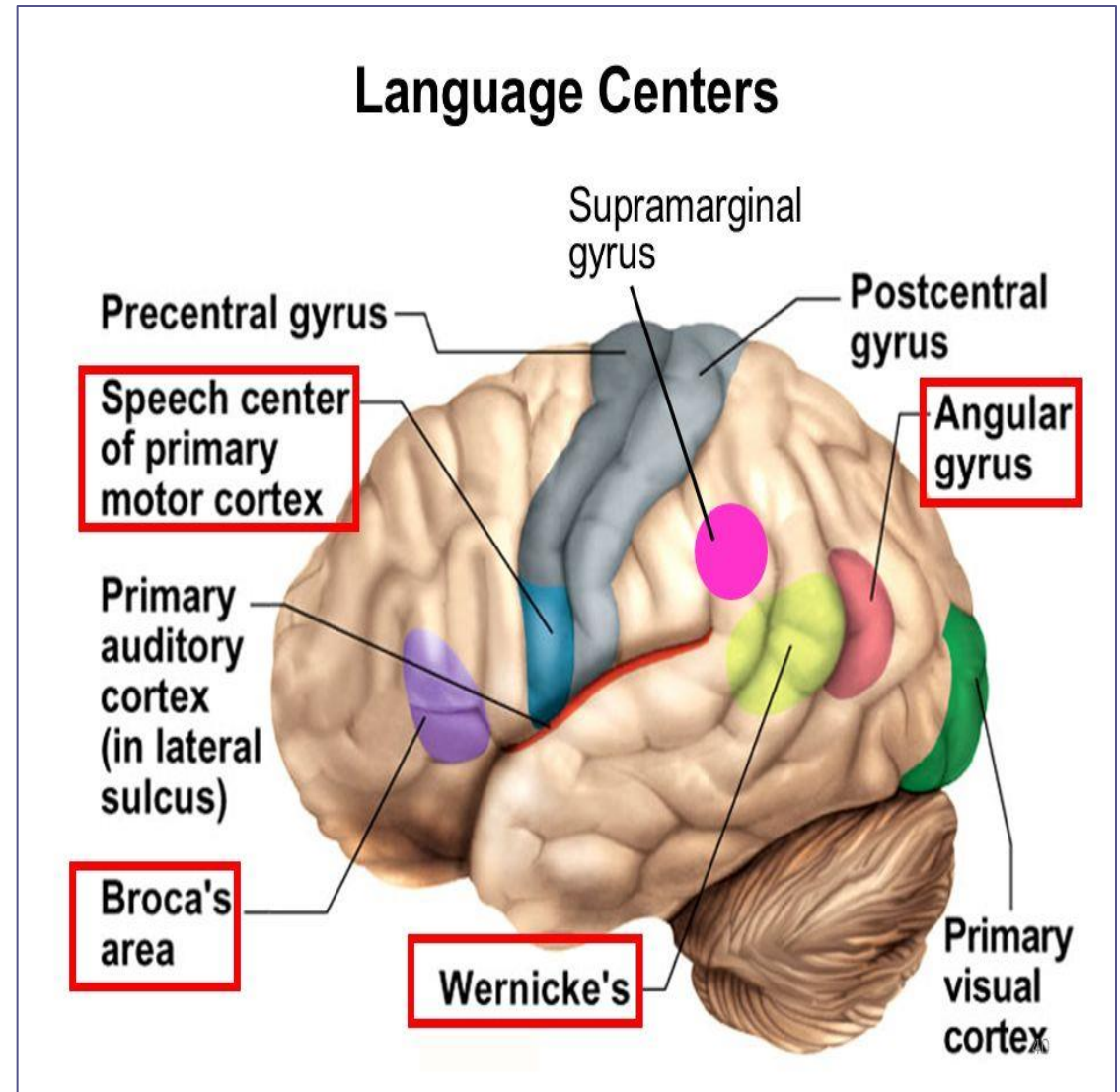
- a. **To speak**, word comprehension originates in Wernicke's area (after receiving information from the auditory and visual cortex) and is sent to **Broca's area** along the arcuate fasciculus (a fiber tract).
- b. **Broca's area** sends information to the motor cortex (precentral gyrus) to direct movement of appropriate muscles of speech.





# Angular Gyrus

- a. Located at the junction between the parietal, occipital, and temporal lobes
- a. Center for integration of sensory information
- a. Damage here also produces aphasias involved in reading and writing

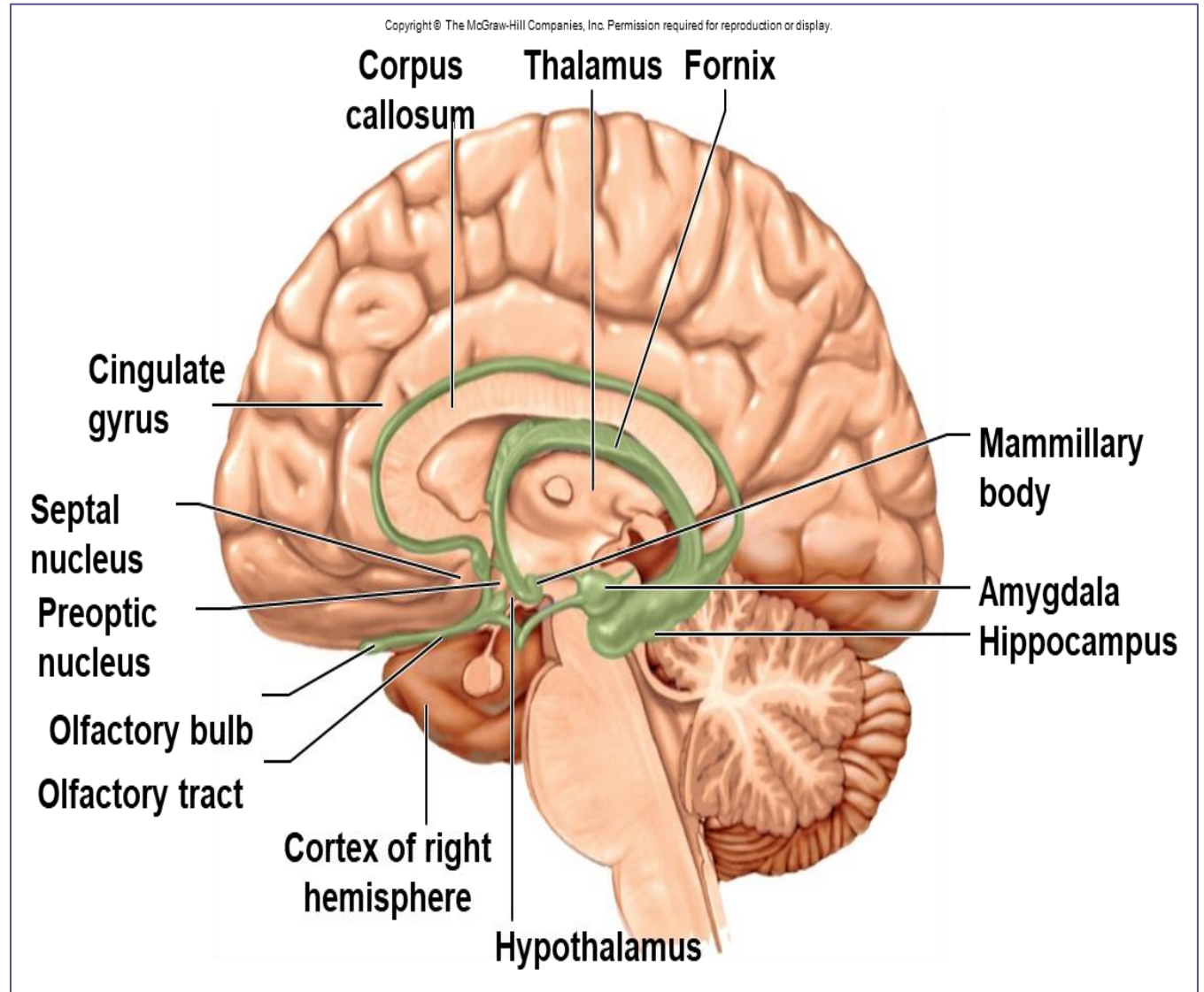


# Aphasias

- a. **Broca's aphasia** results from a damage in Broca's area and involves **slow, poorly articulated speech**; There is **no impairment in understanding**; Weakness in the right arm; Weakness the right side of the face; can understand a sentence but have difficulty repeating it.
- b. **Wernicke's aphasia** involves production of **rapid speech with no meaning**, called **“word salad”**; **Language (spoken and written) comprehension is destroyed**.
- c. **Conduction aphasia**: caused by damage to the arcuate fasciculus; both Broca's and Wernicke's areas are intact; People with Conduction aphasia **are fluent but nonsensical speech as in Wernicke's aphasia**.

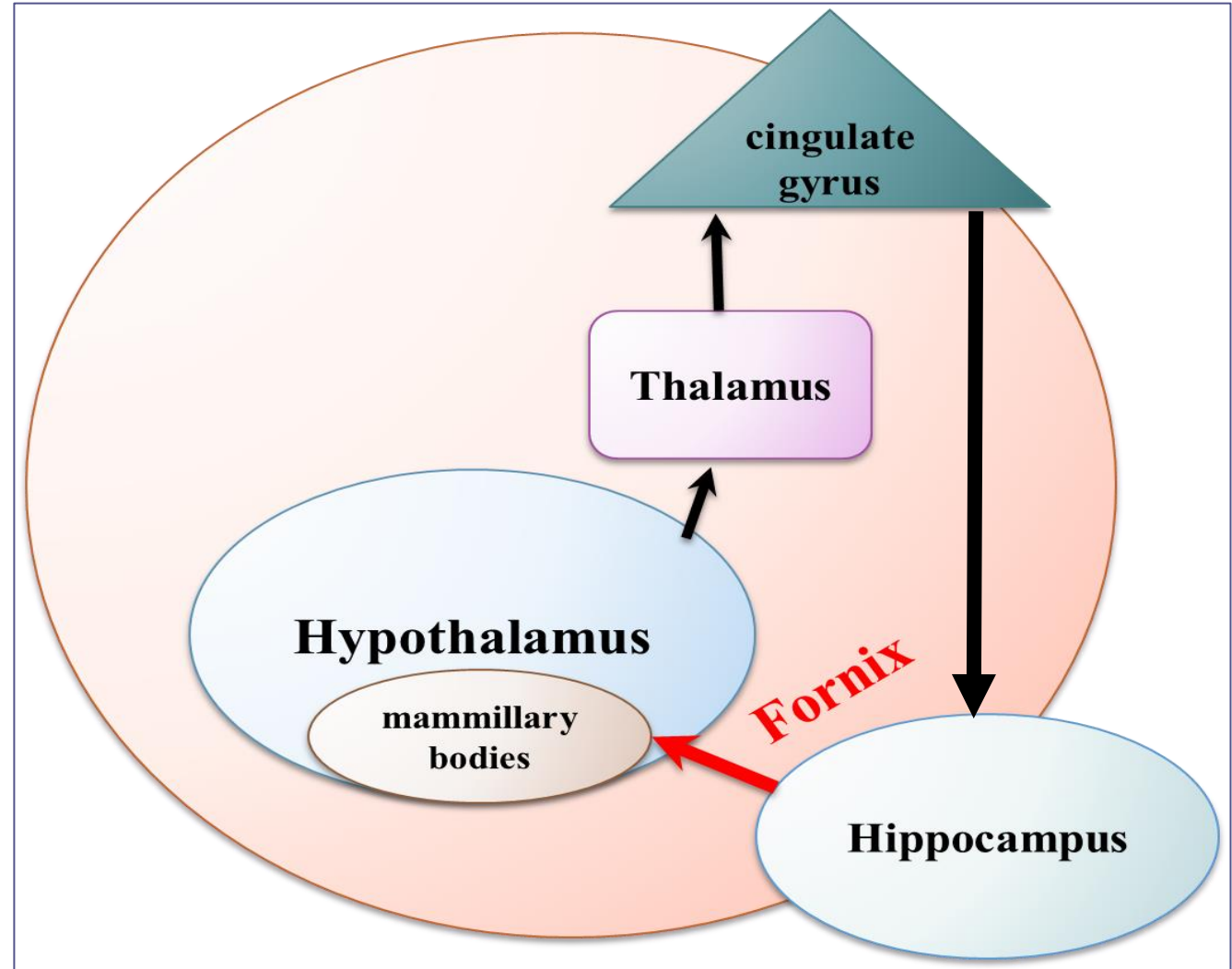
# The Limbic System

- Group of brain regions responsible for **emotional drives (emotional states)**
- A. Areas of the cerebrum included: **cingulate gyrus, amygdala, hippocampus, septal nuclei, anterior insula.**
- B. **The hypothalamus and thalamus** (in the diencephalon) are also part of this system



# The Limbic System (Papez circuit)

- The **fornix** (fiber tract) connects the **hippocampus** to the **mammillary bodies of the hypothalamus**, which sends neurons to the **thalamus**.
- The **thalamus** sends neurons to the **cingulate gyrus**, which sends neurons to the **hippocampus**, completing the circuit.



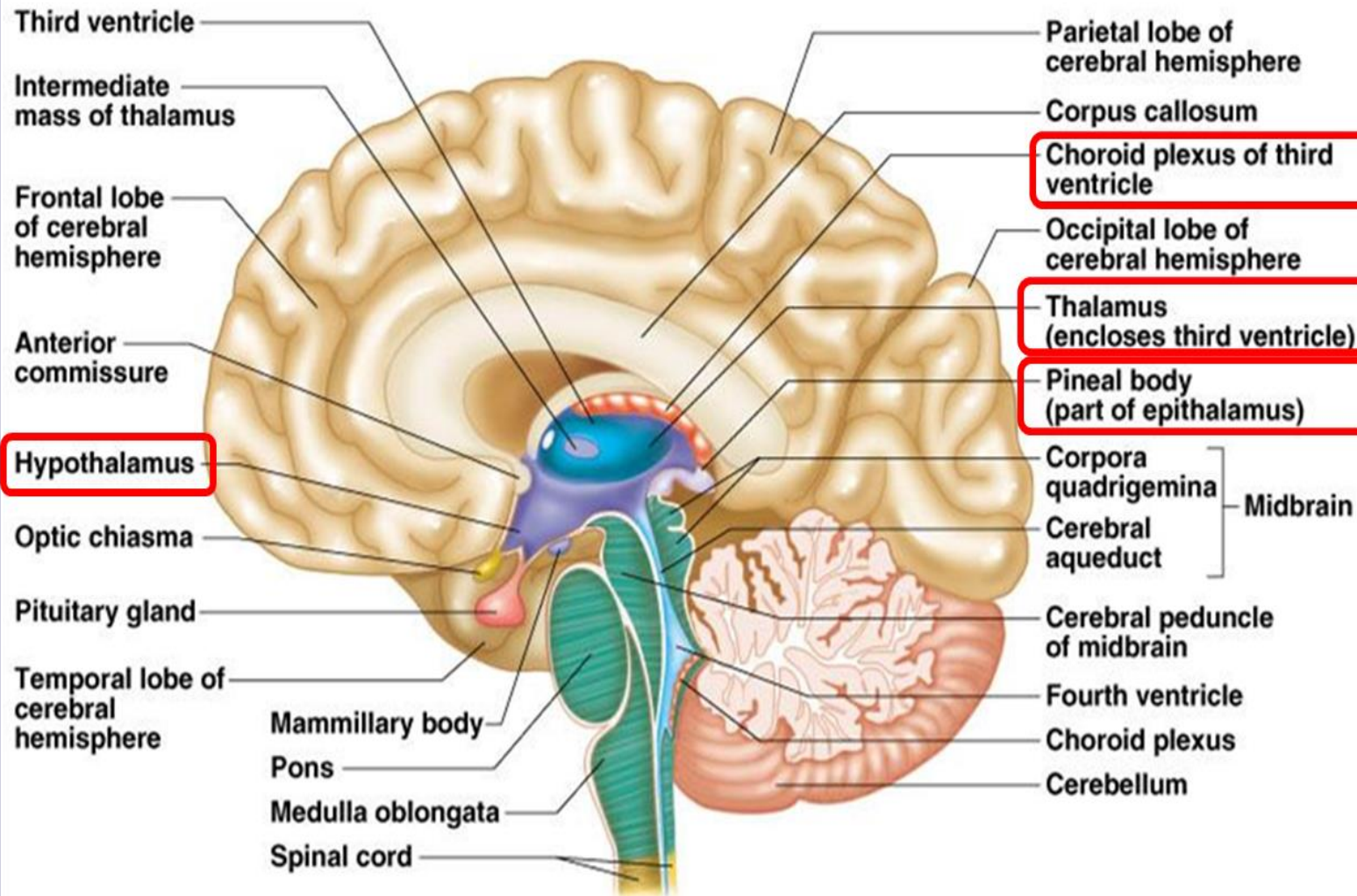
# Limbic System

Emotions controlled by the limbic system:

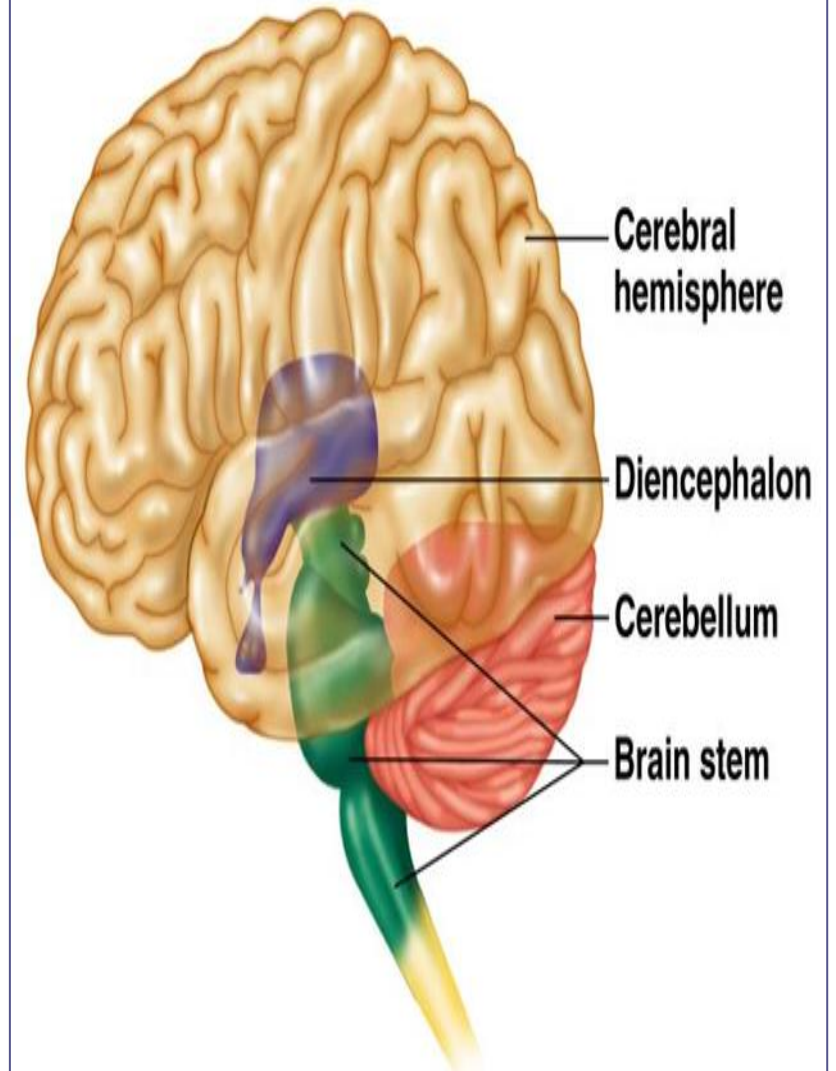
1. **Aggression**: areas in the amygdala and hypothalamus.
2. **Fear**: amygdala and hypothalamus.
3. **Hunger/satiety**: hypothalamus.
4. **Sex drive**: the whole system.
5. **Goal-directed behaviors**: hypothalamus and other regions



# Diencephalon



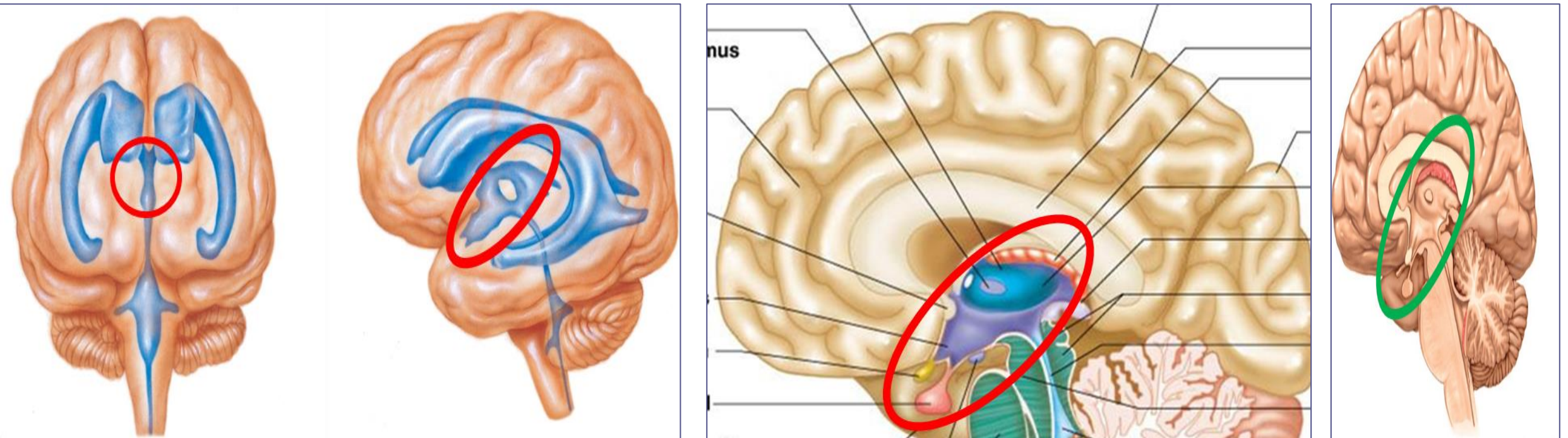
(a)





# Diencephalon

1. It includes the **epithalamus, thalamus, hypothalamus, part of the pituitary gland (posterior pituitary), and the third ventricle.**
2. Surrounded by the **cerebral hemispheres**



# Thalamus

- Forms most of the walls of the **third ventricle**.
- Paired masses of gray matter.
- **Relay center** through which all sensory information, except smell, is passed to the cerebrum.
- The thalamus consists of:
  1. **Geniculate nuclei**
    - A. **lateral geniculate nuclei**: relay visual information from the thalamus to the occipital lobe of the cerebral cortex.
    - B. **medial geniculate nuclei**: relay auditory information from the thalamus to the temporal lobes of the cerebral cortex.
  2. **Intralaminar nuclei** promote a state of arousal from sleep and alertness.

# Epithalamus

- a. The **dorsal** segment of the **diencephalon**
- b. Contains the **choroid plexus** over the **third ventricle** where cerebrospinal fluid is produced
- c. Also contains the **pineal gland**, which secretes the hormone **melatonin** that helps regulate **circadian rhythms**.

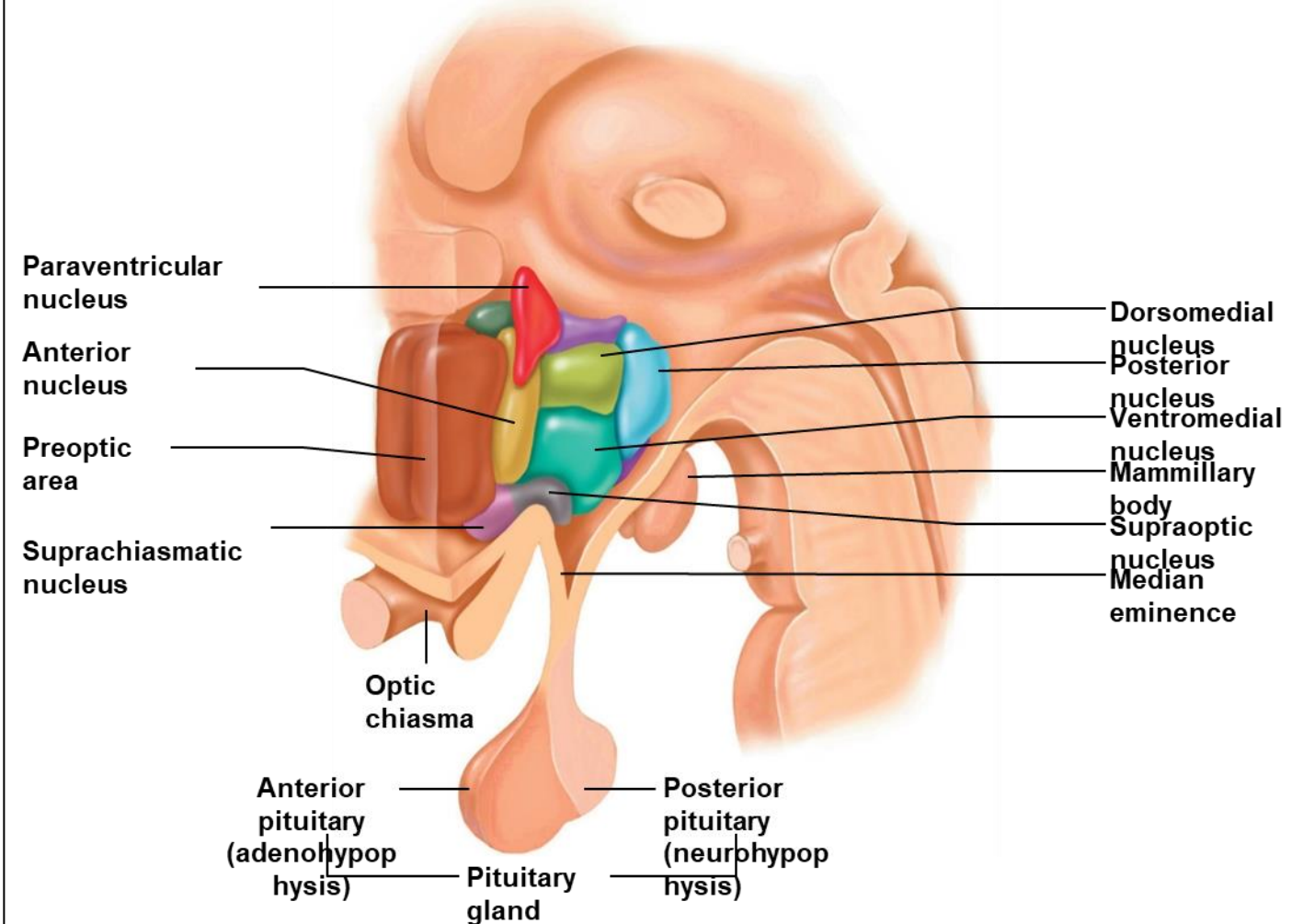
# Hypothalamus

- the most **inferior** portion of the **diencephalon**.
- located **below** the **thalamus**.
- forms the **floor** and part of the **lateral walls** of the **third** ventricle.
- Very important for **maintaining homeostasis** and **regulating the autonomic system**.
- The hypothalamus contains centers for:
  1. **Hunger/satiety** and **thirst**
  2. **Regulation of body temperature**
  3. **Regulation of sleep and wakefulness**
  4. **Sexual arousal** and **performance**
  5. **Emotions of fear, anger, pain, and pleasure**
  6. **Control of the endocrine system**
  7. **Controls hormone secretion from the pituitary gland**

# Regions of the Hypothalamus & Functions

- 1) **Lateral** hypothalamus region: **hunger**
- 2) **Medial** hypothalamus region: **satiety**
- 3) **Preoptic-anterior hypothalamus**: **shivering, hyperventilation, vasodilation, sweating**  
(Hypothalamus contains body's "thermostat" to correct the temperature deviations).
- 4) **Supraoptic**: produces **antidiuretic hormone** and stores it in the posterior pituitary, which helps control urine formation and stimulates thirst.
- 5) **Paraventricular**: produces the hormone **oxytocin**, which stimulates childbirth.

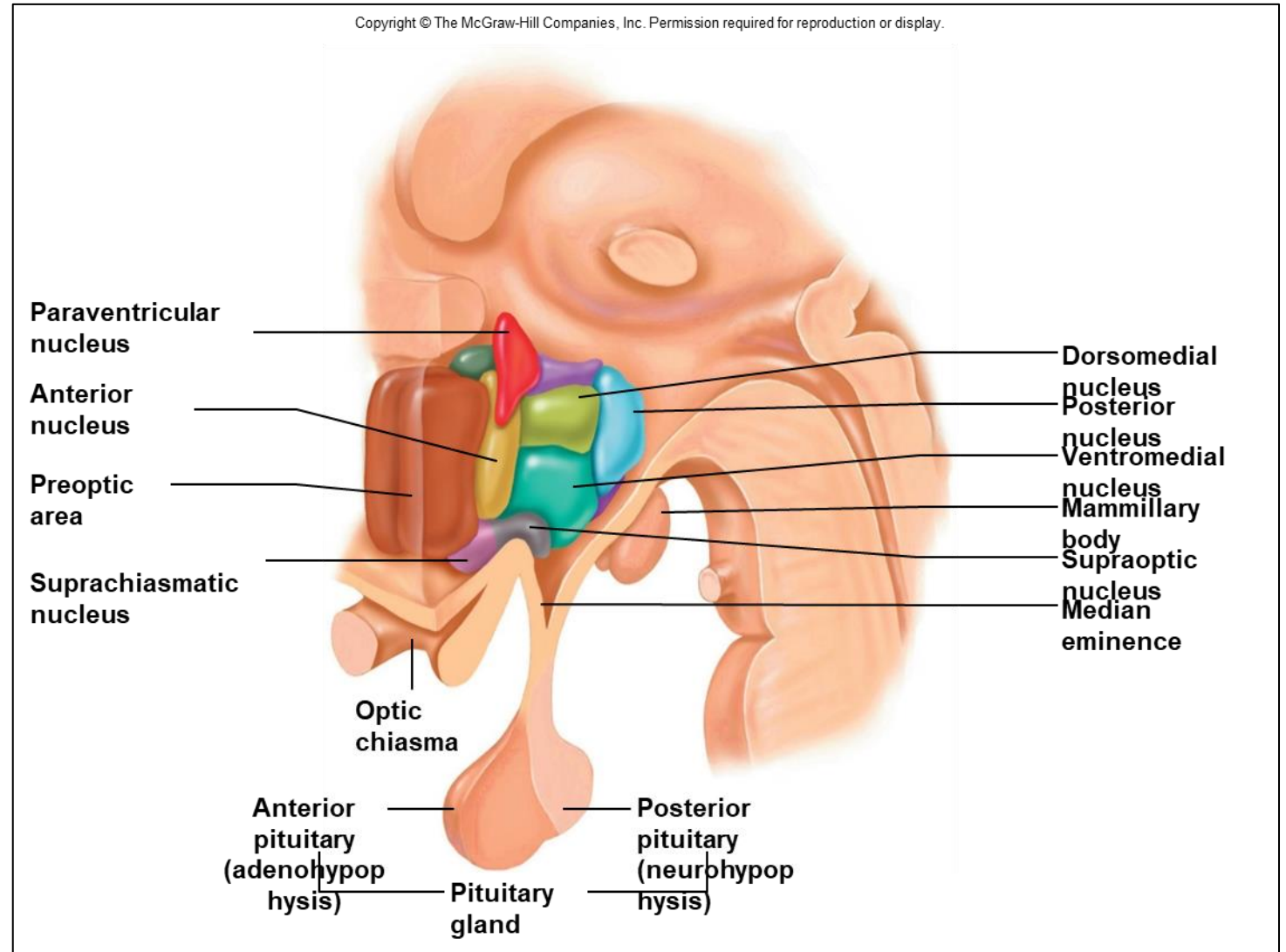
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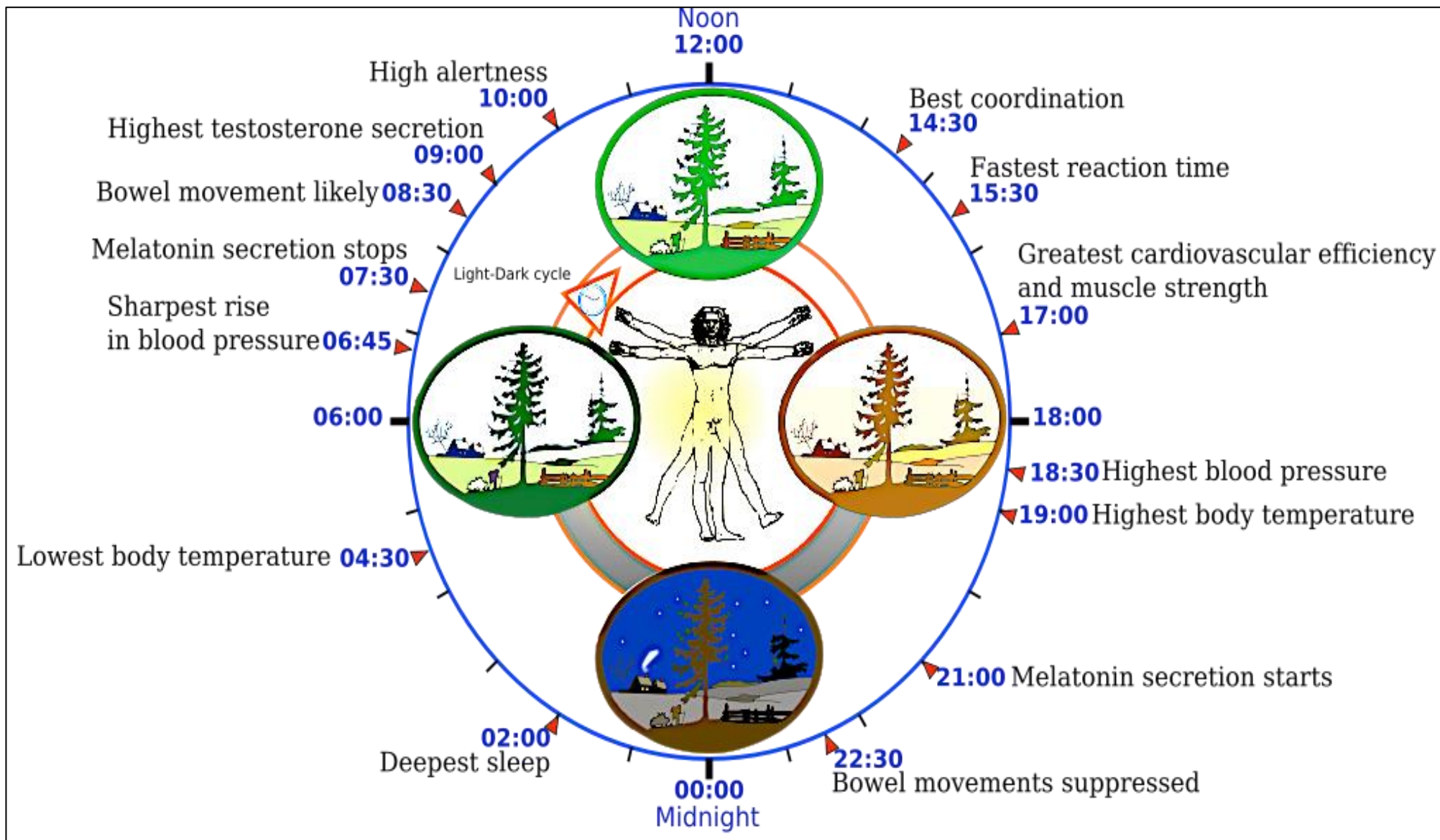


# Regulation of the Pituitary Gland

- 1) **Pituitary gland:** located inferior to the **hypothalamus**.
- 2) **Posterior pituitary** derives from the **diencephalon**.
- 3) **Pituitary gland is connected** to the diencephalon by a stalk.



# Circadian Rhythms



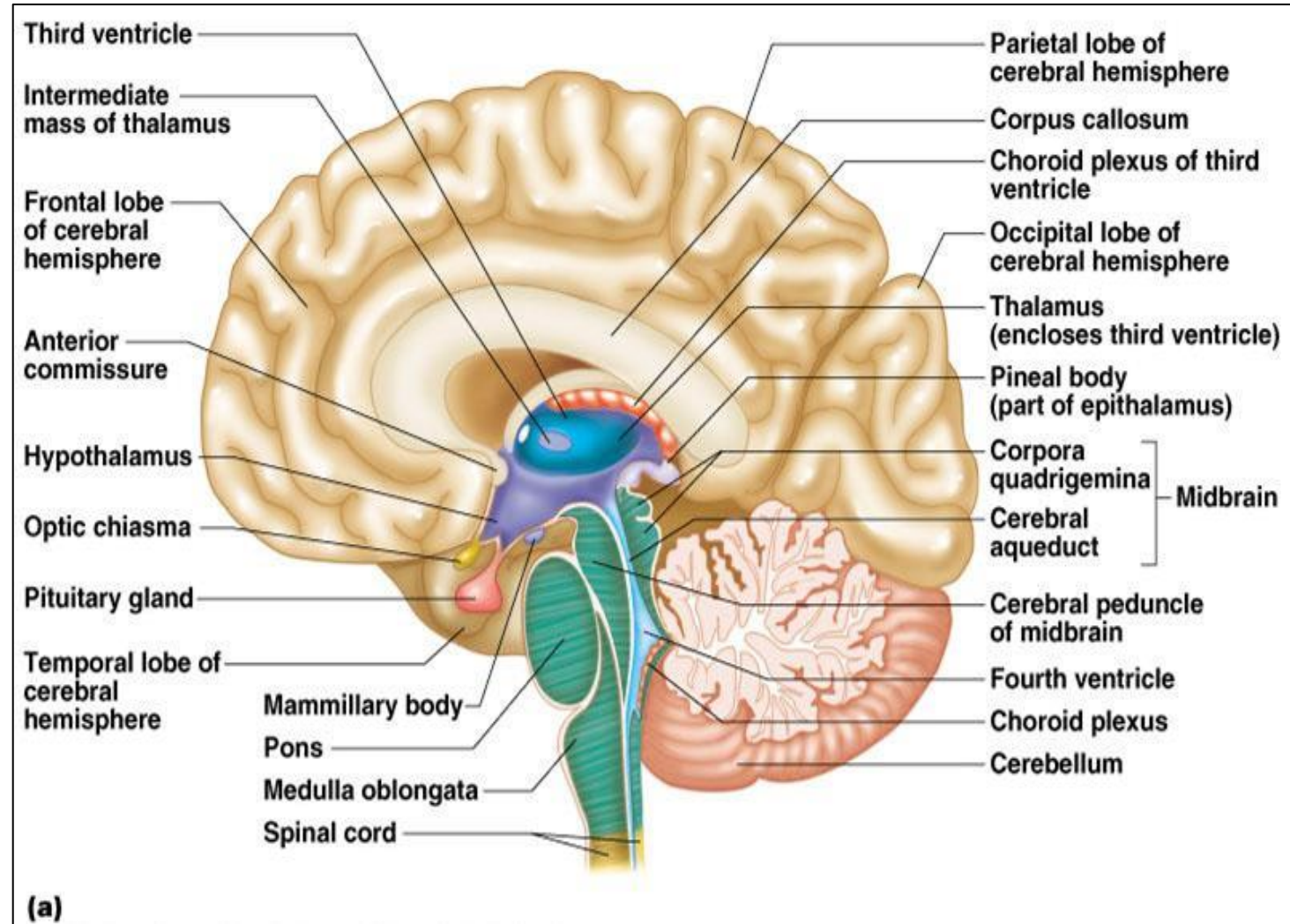
# Regulation of Circadian Rhythms

- 1) **Circadian rhythms**: is any biological process that displays an endogenous, entrainable oscillation of about 24 hours.
- 2) **Circadian clock genes** are found in cells of the **SCN (master clock cells)**, other **(peripheral clock cells)** brain areas, heart, liver, kidneys, skeletal muscle, adipose tissue, and other organs.
- 3) **Suprachiasmatic nuclei (SCN)** are present in the **anterior hypothalamus**: contain about 20,000 “clock cells” with activity that oscillates every 24 hours – main control of circadian rhythms.
- 4) Entrained by information about **day length** via tracts from retinal ganglion cells with the photopigment **melanopsin** by way of **retinohypothalamic tracts**.
- 5) Controls the secretion of **melatonin** from the **pineal gland** which is the major regulator of circadian rhythms; secreted mainly at night
- 6) Neural connections to other brain regions
- 7) Regulation of the pituitary gland

# Midbrain

The midbrain includes:

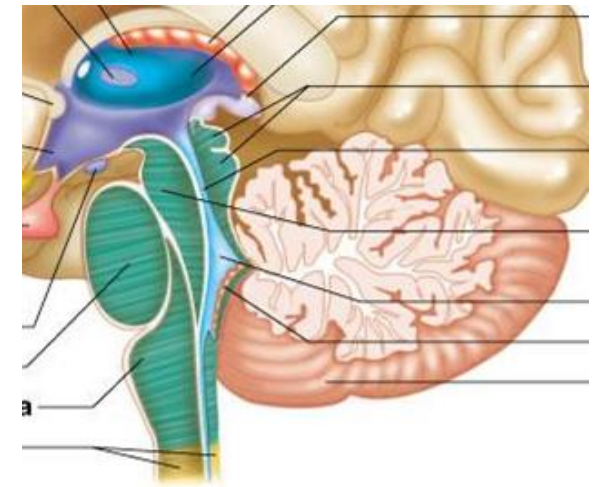
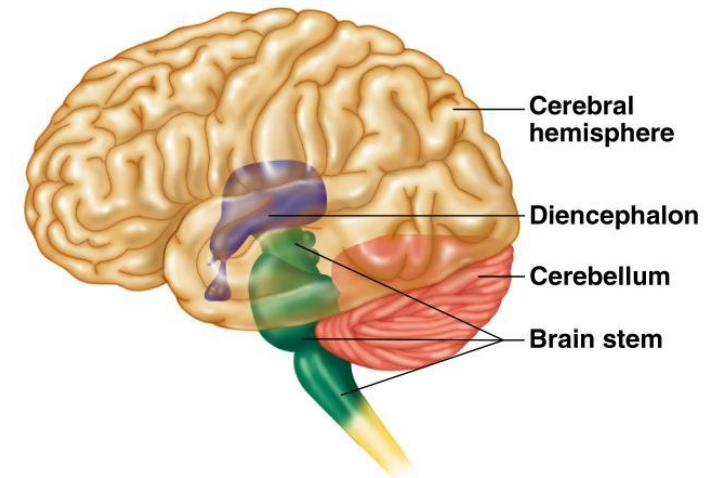
- a. **Corpora quadrigemina** (**four** rounded **elevations** on the dorsal surface)
  - 1) **Superior colliculi**: visual reflexes
  - 2) **Inferior colliculi**: relay center auditory information
- b. **Cerebral peduncles**: ascending and descending tracts
- c. **Red nucleus**: gray matter, connects the cerebrum and cerebellum; involved in motor coordination





# Cerebellum

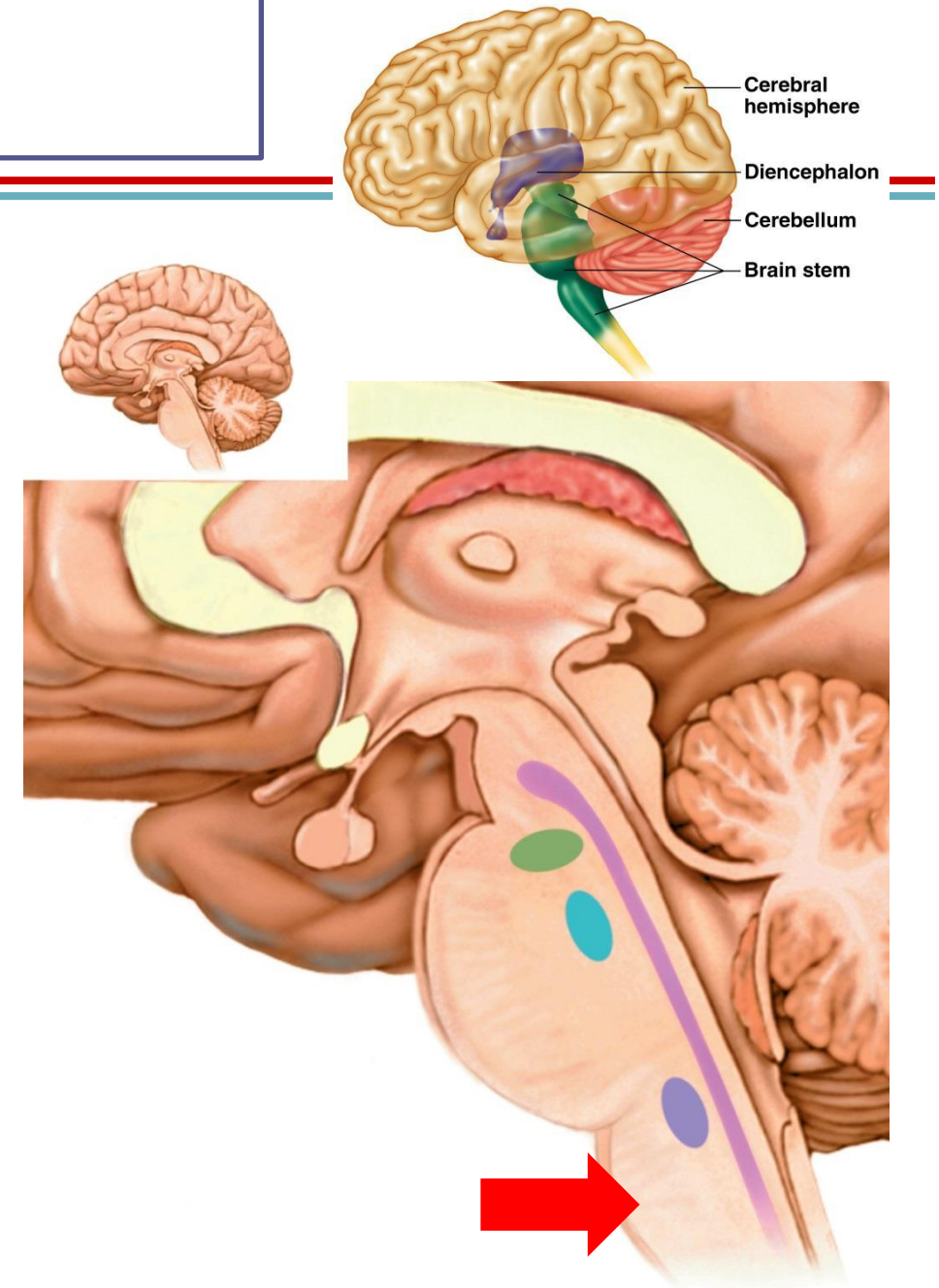
- 1) Second largest brain structure; gray matter outside, white matter inside
- 2) Receives input from proprioceptors in joints, tendons, and muscles
- 3) Works with the basal nuclei and motor cortex to coordinate movement
  - ❖ Fibers from the cerebellum pass through the red nucleus to the thalamus and then to the motor cortex
- 4) The cerebellum is needed for **motor learning** and the **proper timing and force** required to move limbs in a specific task.
- 5) is needed in order to touch your nose with your finger, bring a fork of food to your mouth, or find keys by touch in your pocket or purse.
- 6) The cerebellum influences motor coordination through inhibition on the motor cortex from **Purkinje cells**.
- 7) May have roles in acquisition of sensory data, memory, emotion, and other higher functions





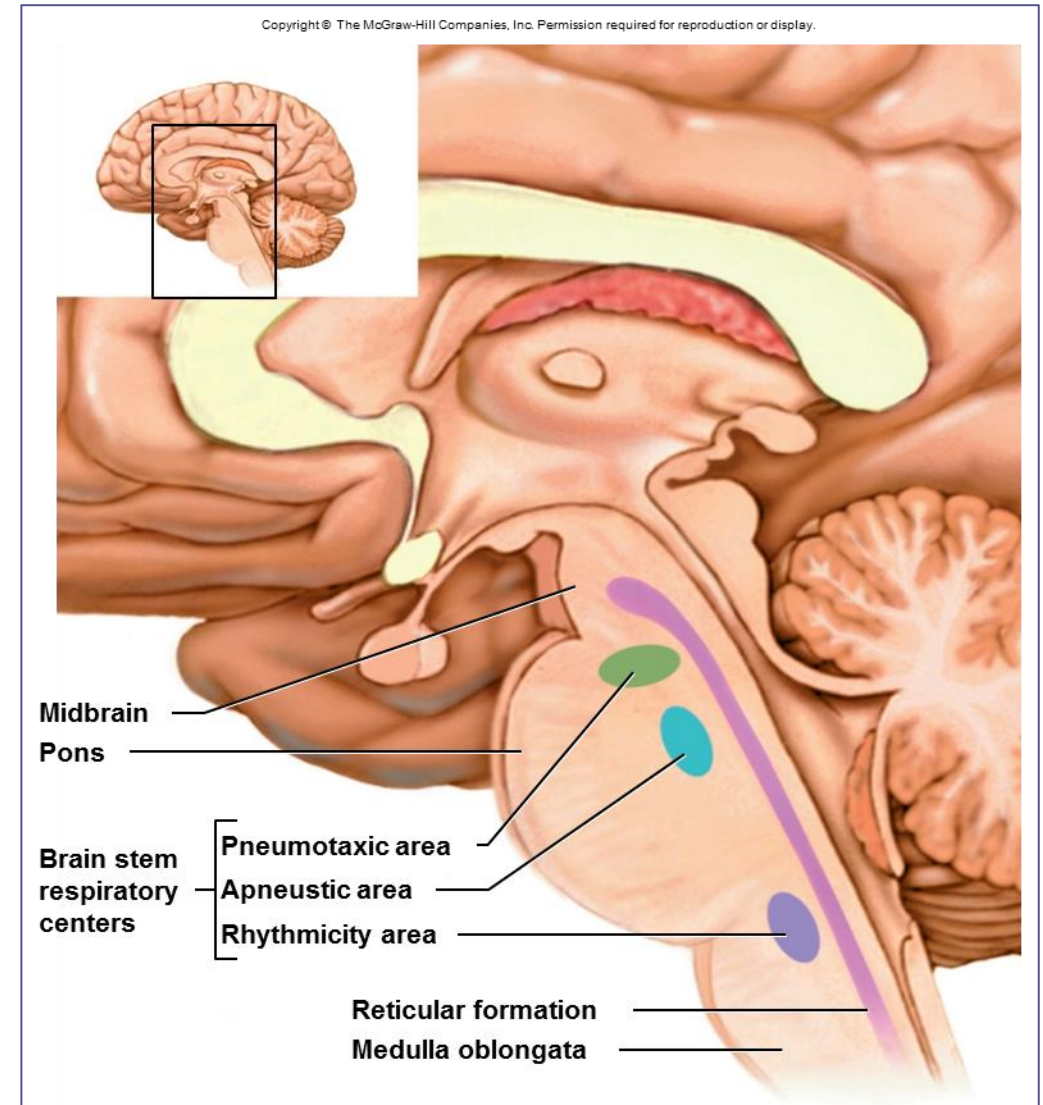
# Medulla oblongata

- All **ascending and descending tracts** between the brain and spinal cord pass through the medulla.
- Tracts **cross sides in the pyramids**.
- **Nuclei of the Cranial nerves VIII, IX, X, XI, and XII** come off the medulla.



# Medulla Oblongata

- Contains **nuclei** required for regulation of **breathing and cardiovascular** response = **vital centers**
- ❖ **Vasomotor center** controls blood vessel diameter (controls the autonomic innervation of blood vessels).
- ❖ **Cardiac control center** controls heart rate (regulates the autonomic nerve control of the heart together with vasomotor center)
- ❖ **Respiratory center** works with areas in the pons to control breathing.



## Ventricles of the CNS

The cavities of the brain are known as **ventricles** (filled with cerebrospinal fluid (CSF)).

The cavity of the spinal cord is called the **central canal**, and is also filled with CSF.

Cerebrospinal fluid is formed by structures called the **choroid plexuses**

*Cerebrospinal fluid has three main functions:*

- 1) CSF **protects** brain and spinal cord from trauma.
- 2) CSF **supplies** nutrients to nervous system tissue.
- 1) CSF **removes** waste products from cerebral metabolism

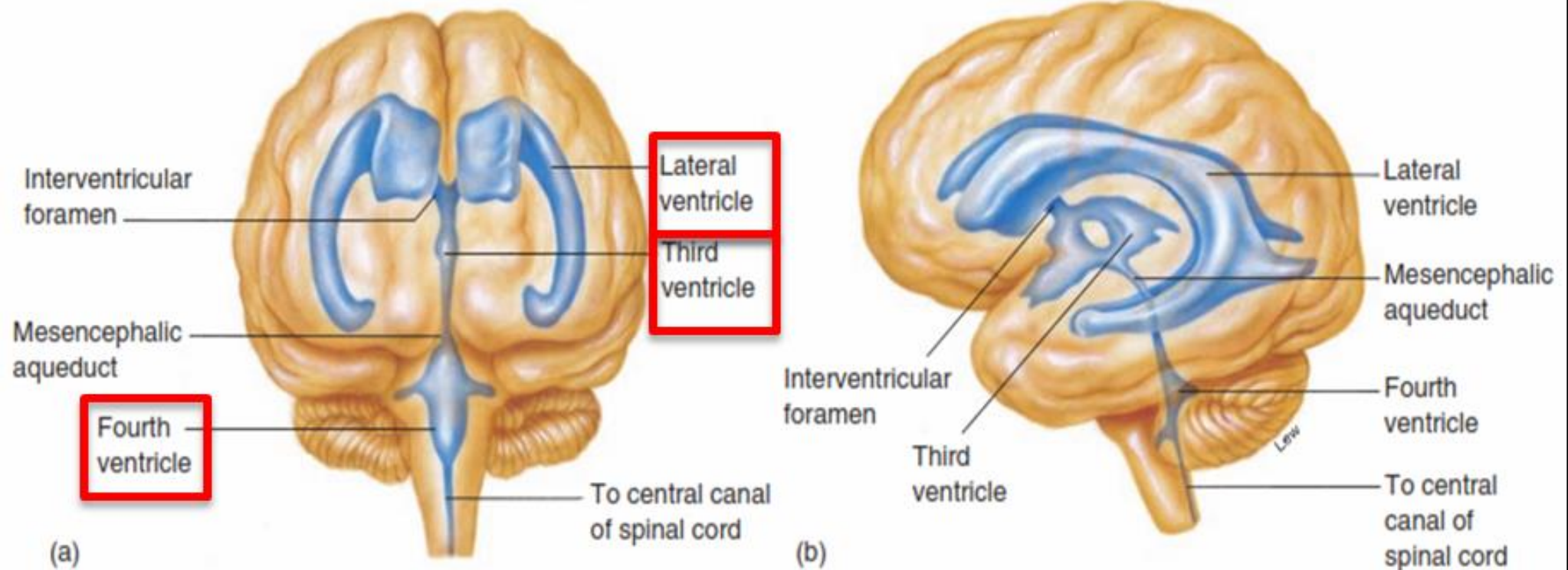
## Ventricles of the CNS

The system comprises four ventricles:

lateral ventricles right and left (one for each hemisphere)

A. Third ventricle

B. Fourth ventricle

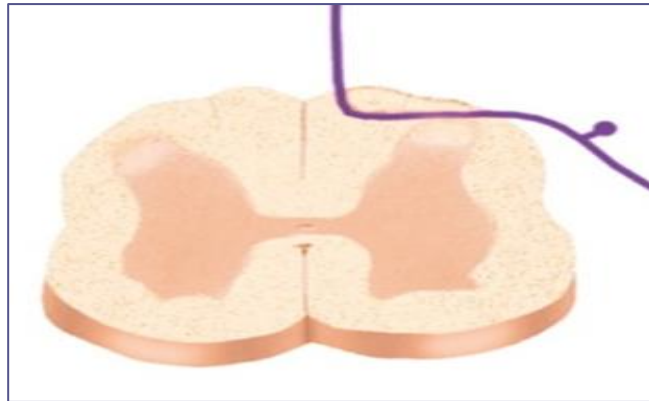


**Figure 8.4** The ventricles of the brain. (a) An anterior view and (b) a lateral view.



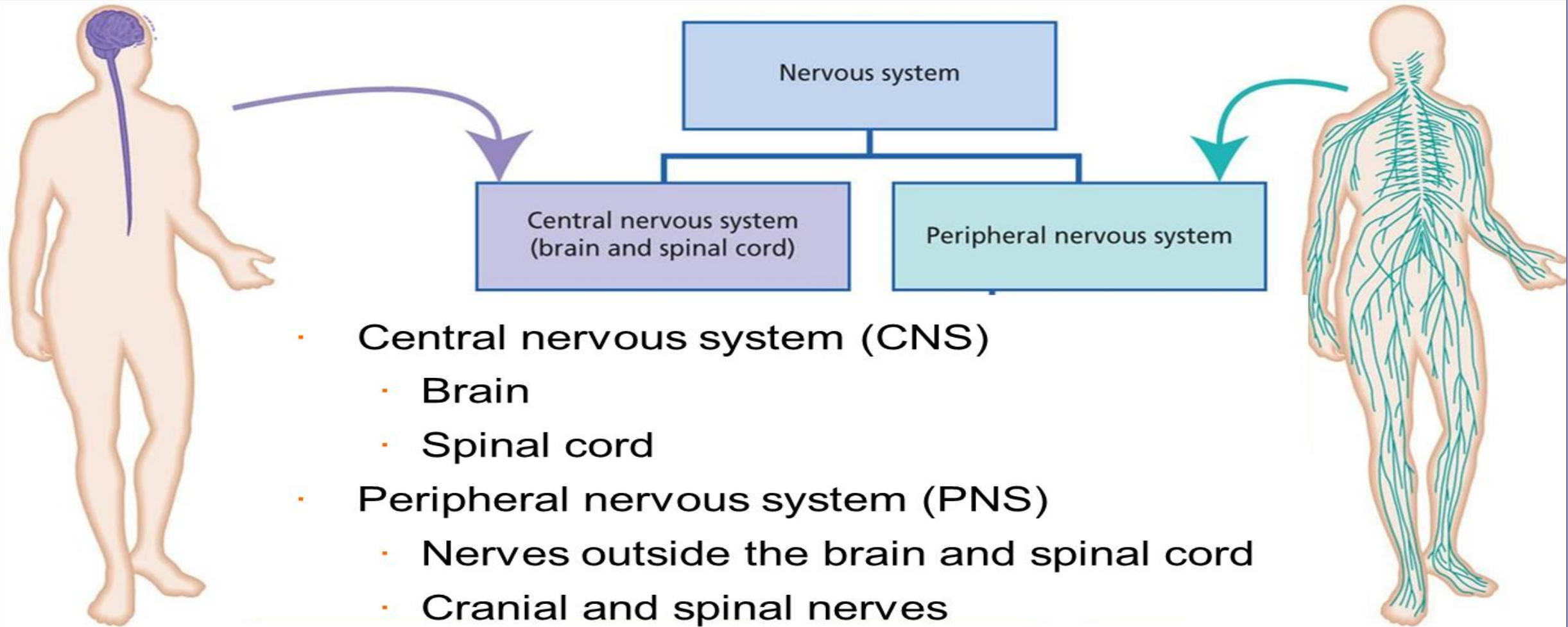
# Spinal Cord Tracts

1. **Spinal cord** acts as a **relay center** for sensory information and motor activities.
2. The spinal cord is composed of **white matter** surrounding a **gray matter** core (arranged in the form of an H)
  - a. The gray matter is arranged with a left and right **dorsal horn** and a left and right **ventral horn**.
  - b. The **white matter** is composed of **ascending** and **descending fiber tracts**.





# Structural Classification of the Nervous System



# Peripheral Nervous System

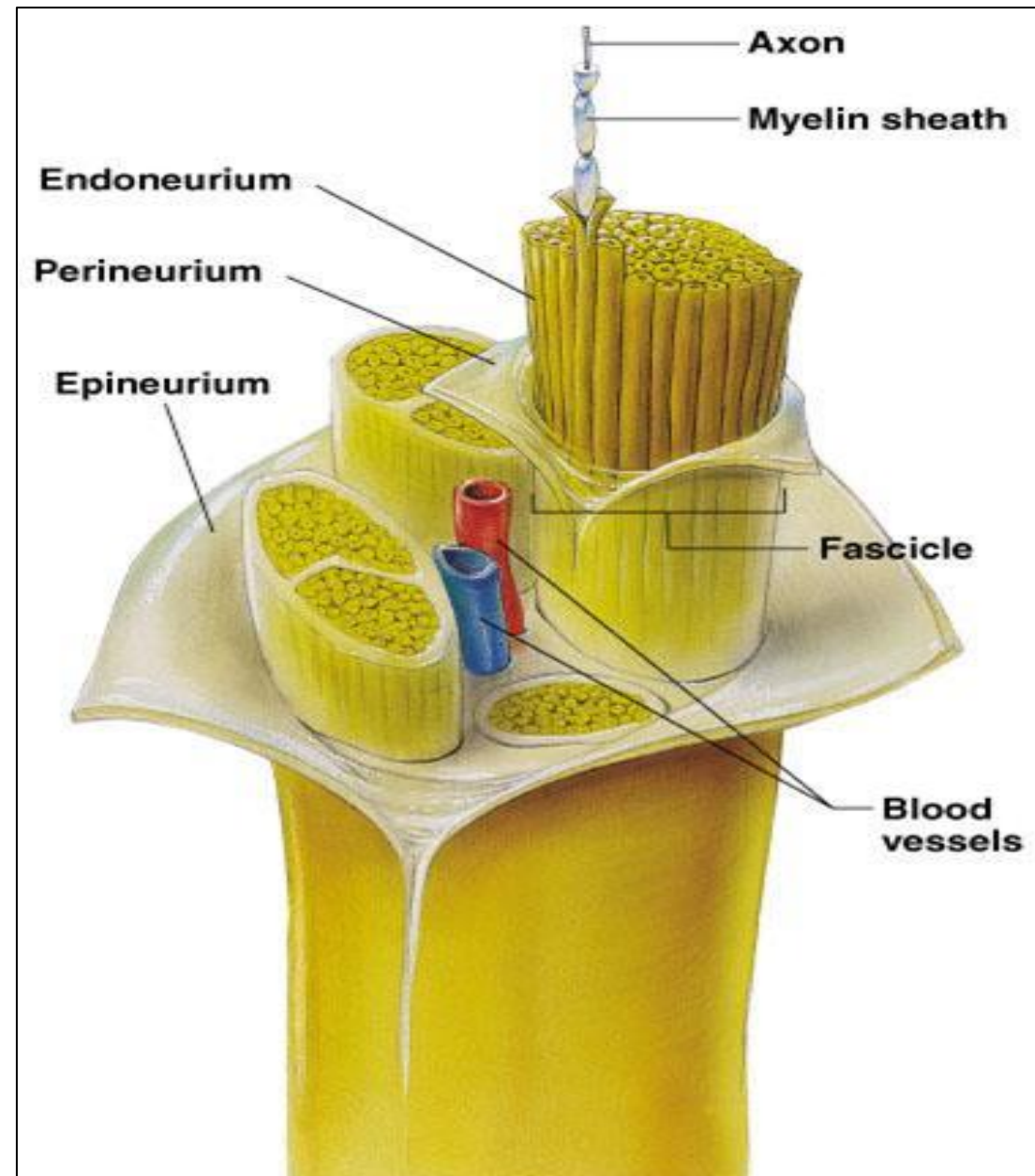
- **Nerves** and **ganglia** **outside** the central nervous system.
- Nerve = **bundle of neuron axons**.
- Neuron axons are **bundled by connective tissue**.
- **Composed of 12 pairs of cranial nerves**.
- **31 pairs of spinal nerves**.

# Structure of a Nerve

**Nerve** – cordlike organ of the PNS consisting of peripheral axons enclosed by connective tissue

## Classification of Nerves

- **Mixed nerves** – both sensory and motor axon bundles
- **Afferent (sensory) nerves** – carry impulses **toward the CNS**
- **Efferent (motor) nerves** – carry impulses **away from the CNS**



# Cranial Nerves

- 12 pairs of nerves that mostly serve the head and neck.
- Numbered in order, front to back.
- Most are mixed nerves, but three are sensory only (I, II and VIII).

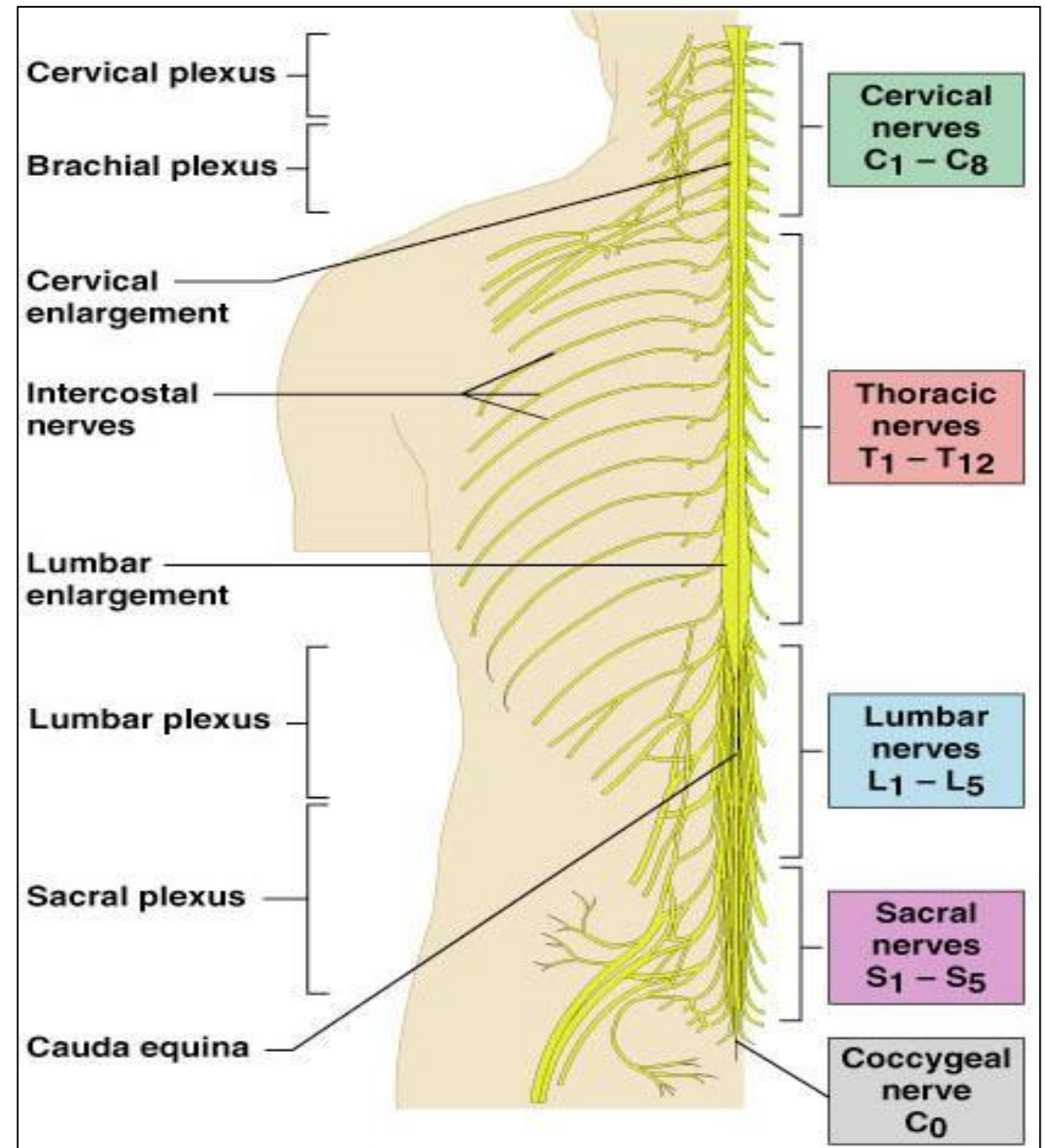
# Cranial Nerves

Number and Name	Composition	Function
I , Olfactory	Sensory	Olfaction
II, Optic	Sensory	Vision
III, Oculomotor	Sensory, Motor	Proprioception and motor impulse to muscles of the eyeball
IV, Trochlear	Sensory, Motor	Proprioception and motor impulse to muscles of the eyeball
V, Trigeminal	Sensory, Motor	Sensory from several parts of the face like the scalp, nasal mucosa, skin of nose and cheek Motor to muscles of mastication (chewing)
VI, Abducens	Sensory, Motor	Proprioception and motor impulse to muscles of the eyeball
VII, Facial	Sensory, Motor	Motor to muscles of facial expression Motor: Secretion of tears and salivation Proprioception from muscles involved in facial expression and sensation from taste buds of the tongue.
VIII, Vestibulocochlear	Sensory	Hearing, Equilibrium (Balance)
IX, Glossopharyngeal	Sensory, Motor	Motor: muscles of pharynx (swallowing), salivation Sensory: taste buds of tongue, Proprioception from muscles of the pharynx
X, Vagus	Sensory, Motor	Motor: pharynx and larynx, regulation of many visceral functions (heart, lung, stomach, intestine,...) Sensory: Proprioception from visceral organs
XI, Accessory	Sensory, Motor	Motor: larynx, muscles of the head, neck and shoulder Sensory: Proprioception from muscles of the head, neck and shoulder
XII, Hypoglossal	Sensory, Motor	Proprioception and motor impulse to muscles of the tongue



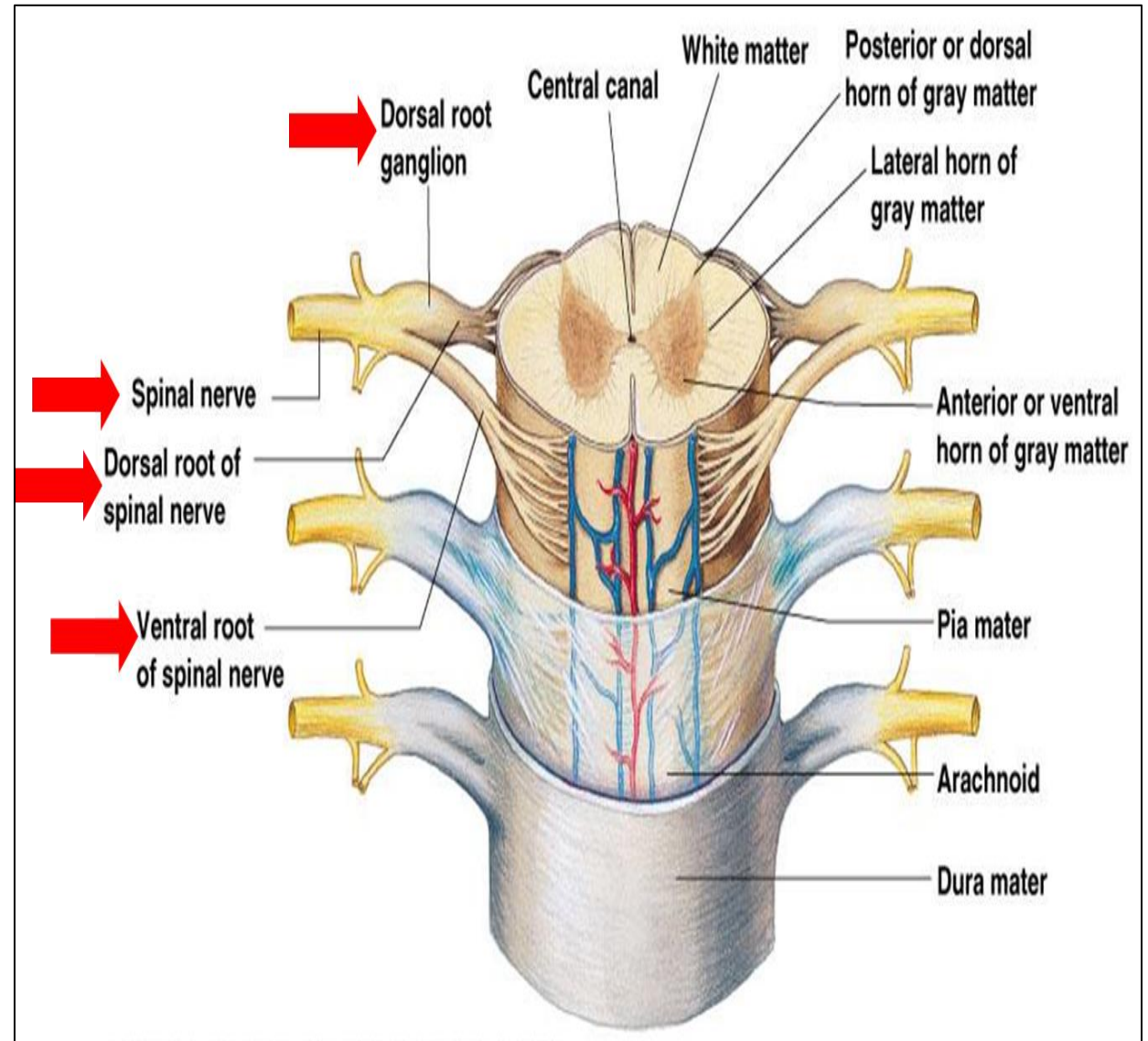
# Spinal Nerves

- They are named for the region of the spinal cord from which they arise.
- 31 pairs



# Spinal Nerves

- All spinal nerves are **mixed nerves**.
- The **cell bodies of the sensory neurons**, whose fibers enter the spinal cord by the **dorsal root**, are found in an enlarged area called the **dorsal root ganglion**.
- The **motor neurons of the somatic nervous system** send their axons through the **ventral root** of the spinal cord.
- The **dorsal and ventral roots fuse** to form the **spinal nerves**.

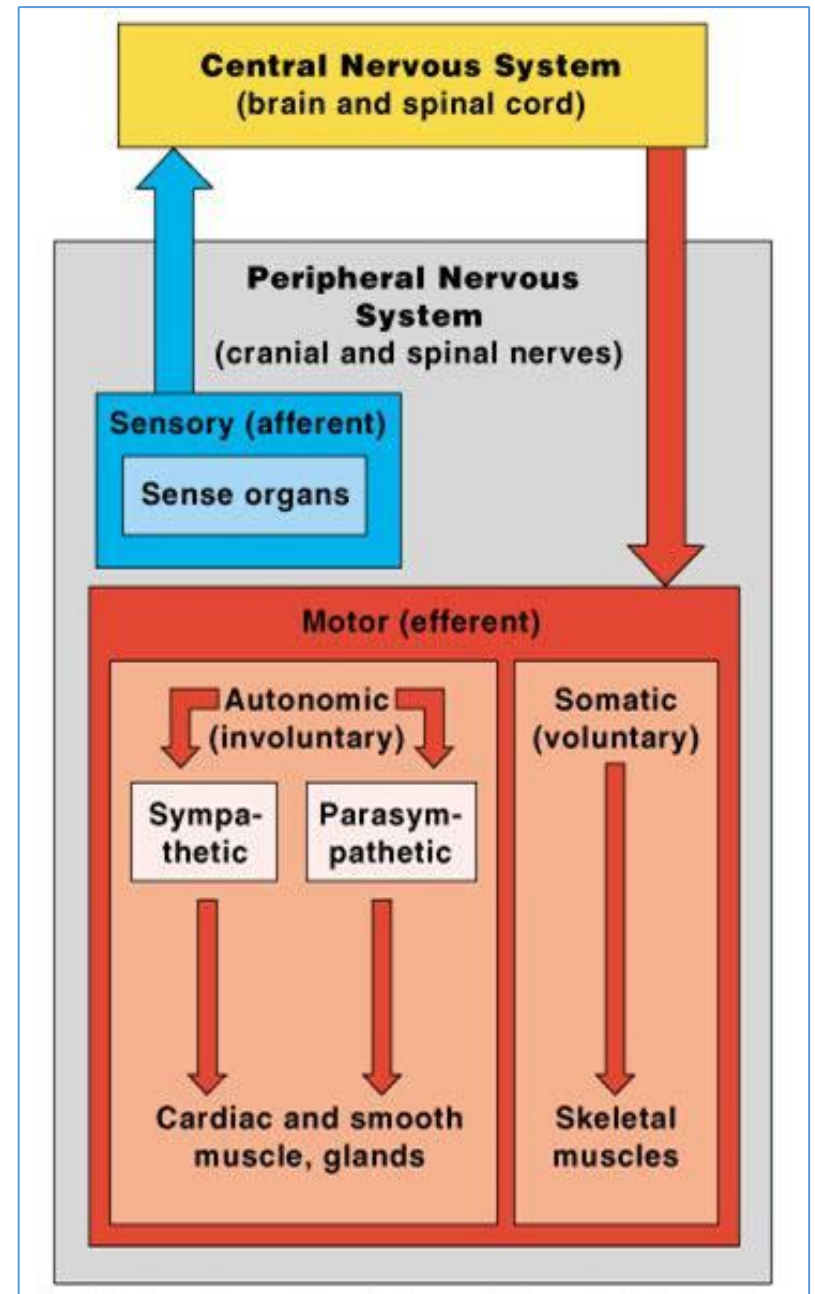


# **Chapter 9**

## **Autonomic Nervous system**

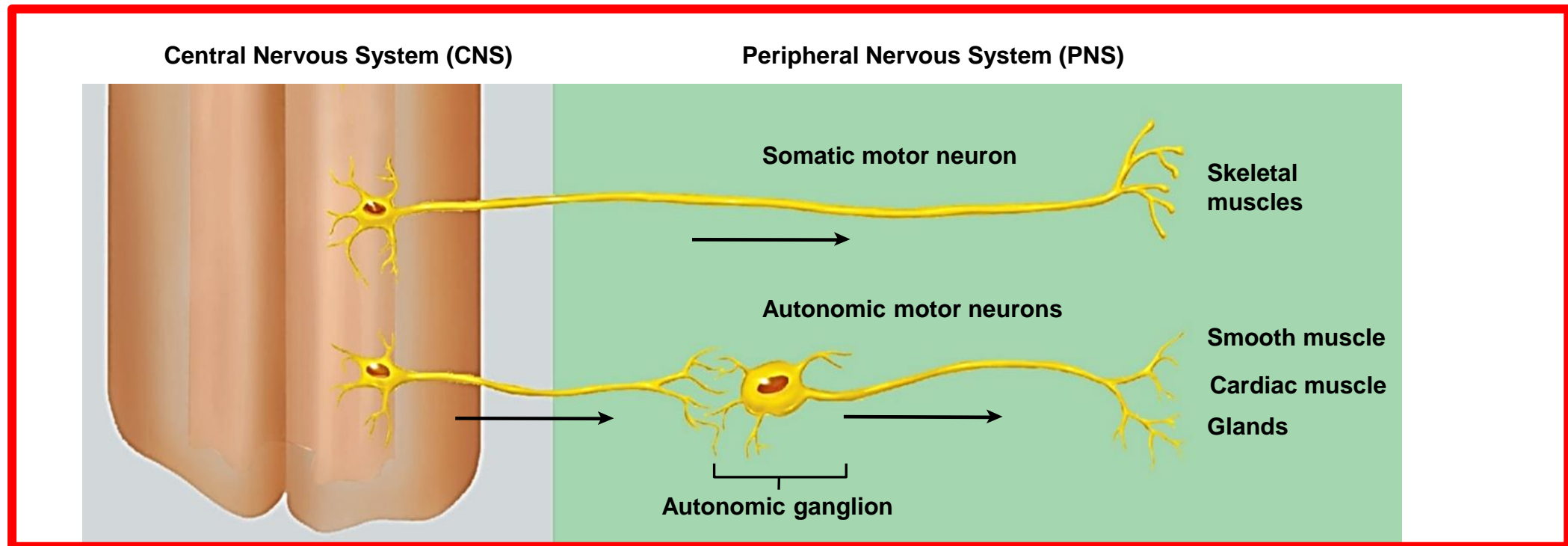
# Autonomic Nervous system

- Innervate organs not under **voluntary** control
- **Effectors include:**
  - A. Cardiac muscle
  - B. Smooth muscle of visceral organs and blood vessels
  - C. Glands
- **Part of the PNS**
- **Neurons are motor, but there are sensory neurons from the viscera** (internal organs and body cavities) for control.



# Differences between somatic and autonomic motor neurons

- Somatic motor neurons have cell bodies in the CNS (brainstem and spinal cord) and just one neuron traveling from CNS to effector.
- The autonomic motor system has two sets of neurons in the PNS.

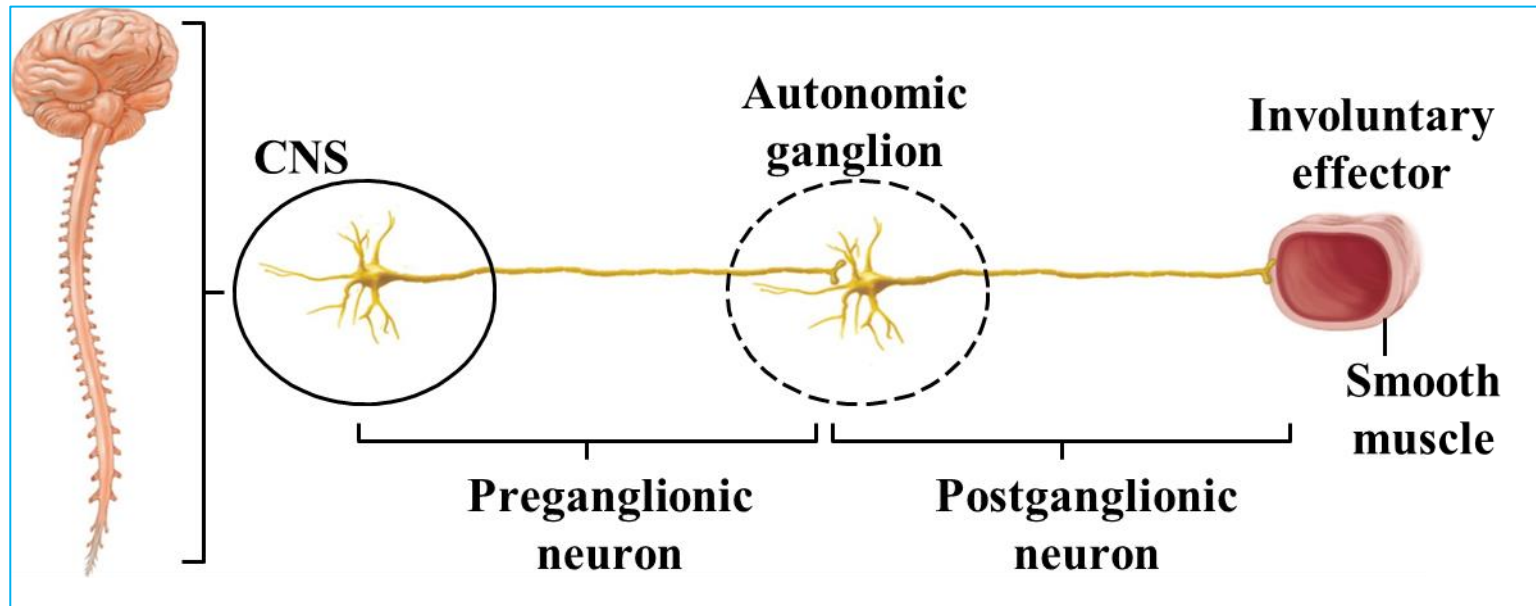




# The ANS has pre- and postganglionic neurons

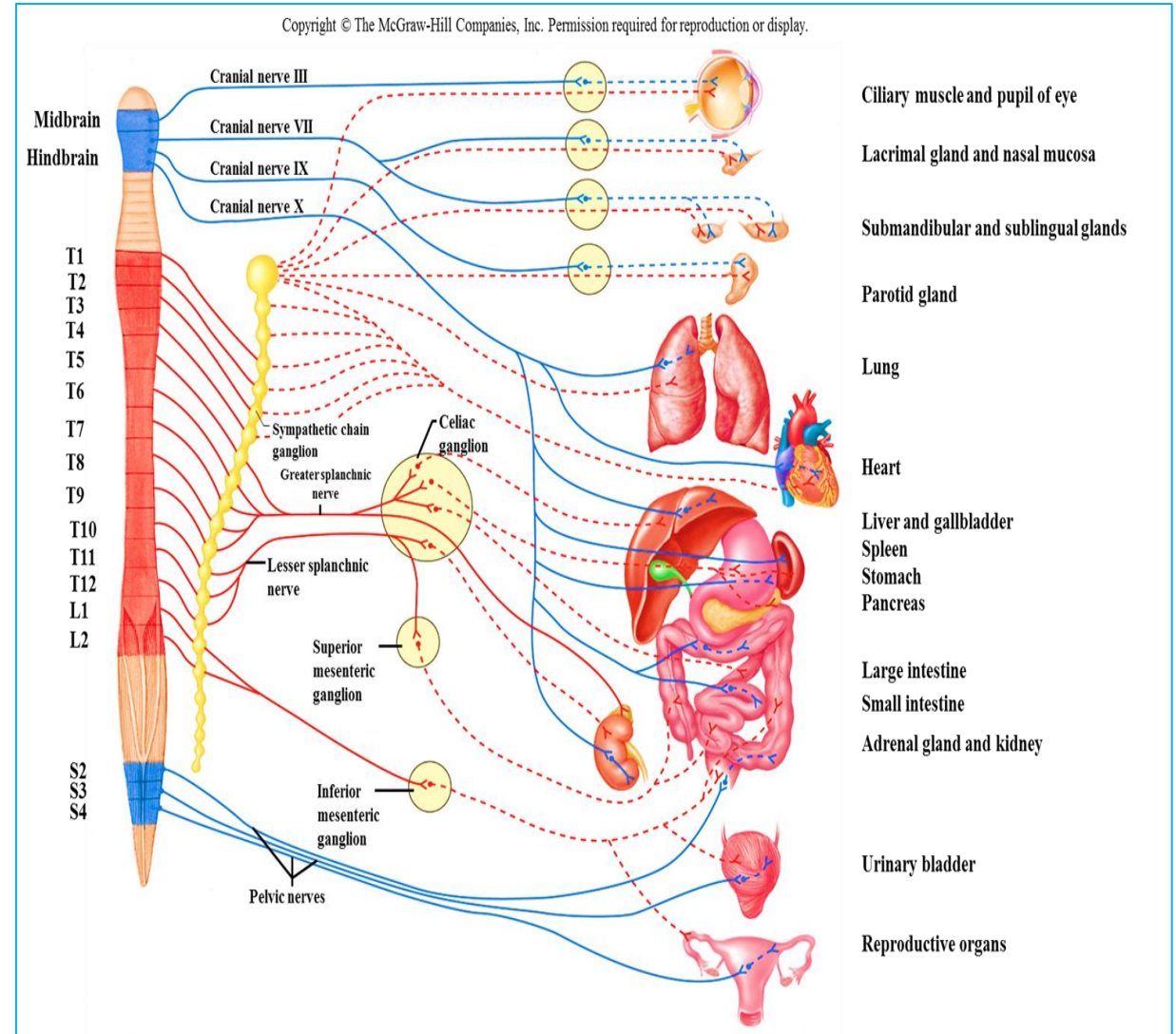
The autonomic motor system has two sets of neurons in the PNS.

1. **The first has cell bodies in** (originates in) the **brainstem or spinal cord** and synapses in an autonomic ganglion (**Preganglionic neuron**).
2. **The second has cell bodies in** (originates in) the ganglion and synapses on the effector (**Postganglionic neuron**).



# The ANS has pre- and postganglionic neurons

- A. **Preganglionic neurons:** originate in the brainstem (midbrain or hindbrain) or from the thoracic, lumbar, or sacral spinal cord.
- B. **Postganglionic neurons:** originate in ganglion.
- C. **Autonomic ganglia** are located in the head, neck, and abdomen as well as in chains along either side of the spinal cord.



## Differences between somatic and autonomic motor neurons: neurotransmitters

- Somatic motor neurons release only **acetylcholine** which is always excitatory.
- Autonomic neurons release mainly **acetylcholine** and **norepinephrine** but may be excitatory or inhibitory

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**Table 9.1 | Comparison of the Somatic Motor System and the Autonomic Motor System**

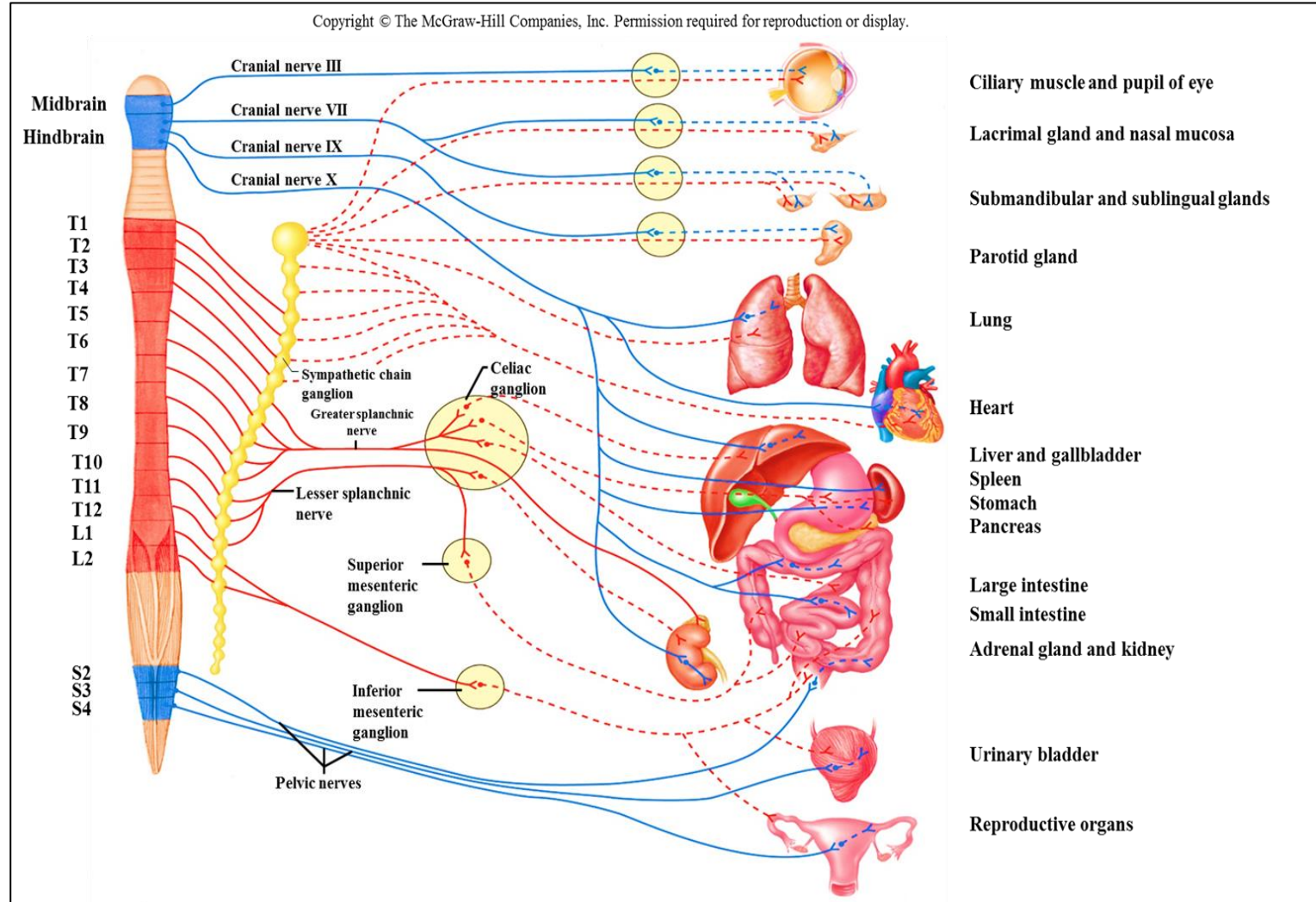
Feature	Somatic Motor	Autonomic Motor
Effector organs	Skeletal muscles	Cardiac muscle, smooth muscle, and glands
Presence of ganglia	No ganglia	Cell bodies of postganglionic autonomic fibers located in paravertebral, prevertebral (collateral), and terminal ganglia
Number of neurons from CNS to effector	One	Two
Type of neuromuscular junction	Specialized motor end plate	No specialization of postsynaptic membrane; all areas of smooth muscle cells contain receptor proteins for neurotransmitters
Effect of nerve impulse on muscle	Excitatory only	Either excitatory or inhibitory
Type of nerve fibers	Fast-conducting, thick (9–13 $\mu$ m), and myelinated	Slow-conducting; preganglionic fibers lightly myelinated but thin (3 $\mu$ m); postganglionic fibers unmyelinated and very thin (about 1.0 $\mu$ m)



# Divisions of the Autonomic Nervous System

## 1. Sympathetic Division

- **Preganglionic neurons** come from the thoracic and lumbar regions of the spinal cord.
- **Sympathetic division** also called the thoracolumbar division
- **Preganglionic neurons synapse** in sympathetic ganglia that run parallel to the spinal cord.
  - A. These are called the **paravertebral ganglia**.
  - B. These ganglia are connected, forming a **sympathetic chain of ganglia**.

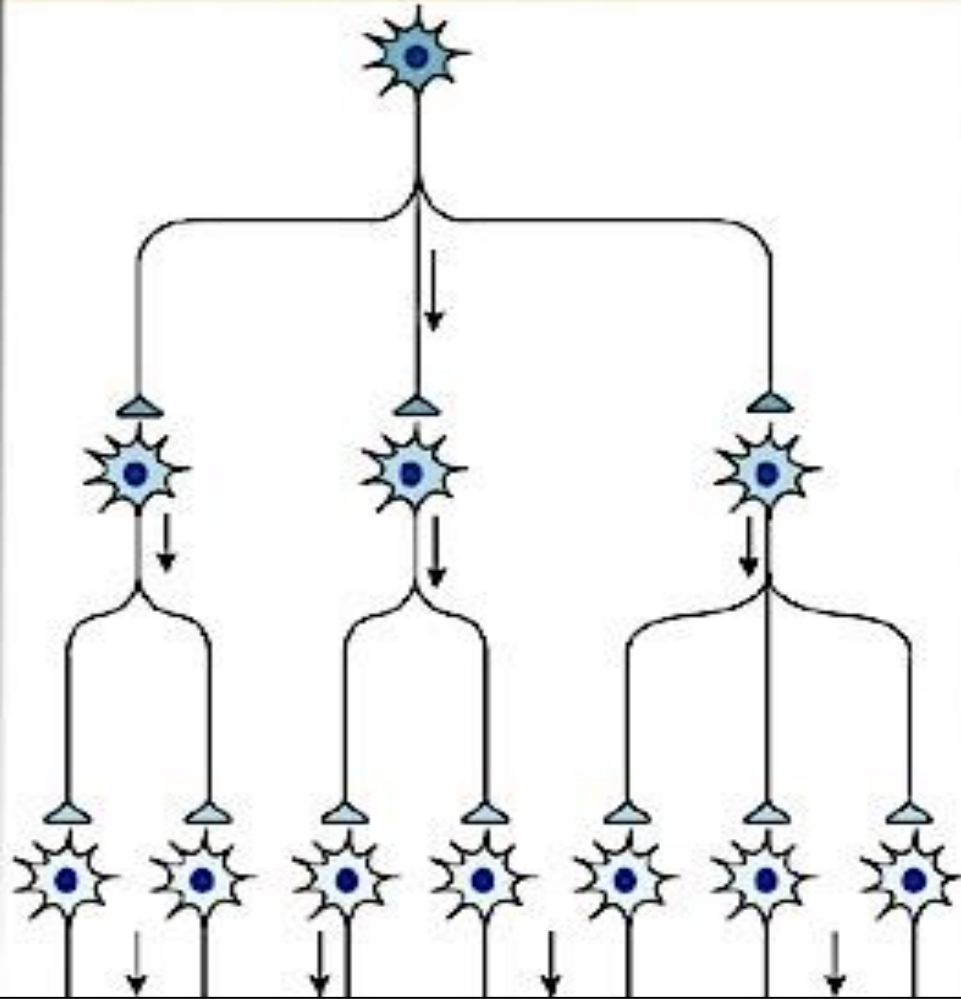


# Convergence and Divergence

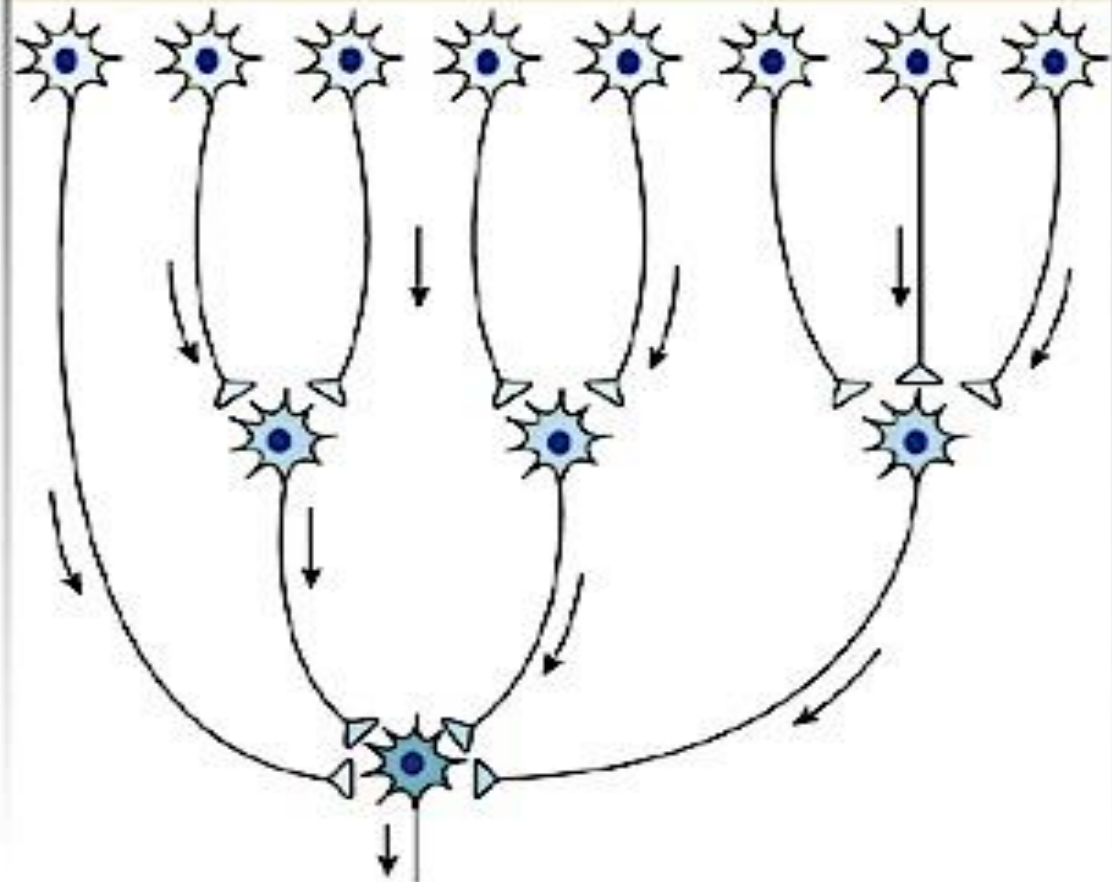
- Because preganglionic neurons can branch and synapse in ganglia at any level, **there is**:
  - 1) **Divergence**: One preganglionic neuron synapses on several postganglionic neurons at different levels.
  - 2) **Convergence**: Several preganglionic neurons at different levels synapse on one postganglionic neuron.
- Allows the sympathetic division to act as a single unit through **mass activation** and to be tonically active.



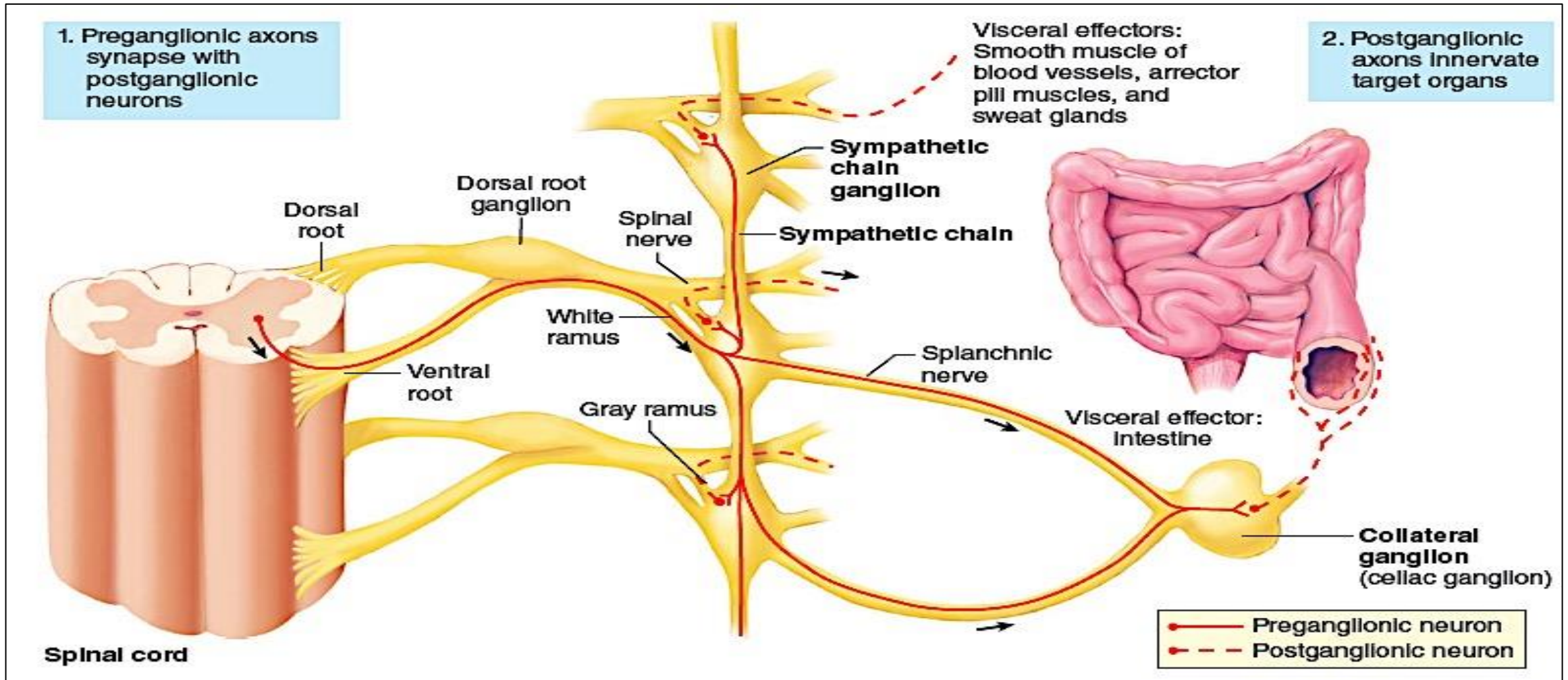
**(a) In a divergent pathway, one presynaptic neuron branches to affect a larger number of postsynaptic neurons.**



**(b) In a convergent pathway, many presynaptic neurons converge to influence a smaller number of postsynaptic neurons.**



# Sympathetic Neuron Pathways



# Parasympathetic Division

- Preganglionic neurons come from the **brain** or **sacral region of the spinal cord**.
  - A. Also called the **craniosacral** division.
  - B. They synapse on ganglia located near or in effector organs; called **terminal ganglia**.
  - C. Terminal ganglia **supply very short** postganglionic neurons to the effectors.

# Functions of the Autonomic Nervous System: General functions

- **Sympathetic Functions:**

- A. The sympathetic division activates the body for “fight or flight” through the release of **norepinephrine** from postganglionic neurons and the secretion of **epinephrine** from the adrenal medulla.
- B. Prepares the body for intense physical activity in emergencies by increasing heart rate and blood glucose levels and by diverting blood to skeletal muscles
- C. Tonically **regulates heart, blood vessels, and other organs.**

- **Parasympathetic Functions**

- A. The parasympathetic division is antagonistic to the sympathetic division.
- B. Allows the body to “rest and digest” through the release of **ACh** from postganglionic neurons.
- C. Slows **heart rate, and increases digestive activities**

# Adrenergic & Cholinergic Synaptic Transmission

## A. Cholinergic Synaptic Transmission

1. **Acetylcholine (ACh)** is the neurotransmitter used by all preganglionic neurons (sympathetic and parasympathetic).
2. It is also the neurotransmitter released from **most** parasympathetic postganglionic neurons.
3. **Some** sympathetic postganglionic neurons (those that innervate sweat glands and skeletal muscle blood vessels) release ACh.
4. These synapses are called **cholinergic**.

## B. Adrenergic Synaptic Transmission

1. **Norepinephrine** is the neurotransmitter released by **most** sympathetic postganglionic neurons.
2. These synapses are called **adrenergic**.



# Response to Adrenergic Stimulation

- Can be epinephrine in the blood or norepinephrine from sympathetic nerves
  - Can stimulate or inhibit, depending on receptors.
- A. **Stimulation**: heart, dilatory muscles of the iris, smooth muscles of many blood vessels (causes vessel constriction).
- B. **Inhibition**: Bronchioles in lungs, other blood vessels; inhibits contraction and causes dilation of these structures

# $\alpha$ and $\beta$ Adrenergic Receptors

1. Two types of  $\alpha$  (alpha) -  $\alpha_1$  and  $\alpha_2$
2. Three types of  $\beta$  (beta) -  $\beta_1$ ,  $\beta_2$  and  $\beta_3$
3. All act using G-proteins and **second messenger systems**.
  - A.  $\beta$  receptors use cAMP.
  - B.  $\alpha$  receptors use a  $\text{Ca}^{2+}$  second messenger system.
4. Alpha receptors are more sensitive to norepinephrine
5. Beta receptors are more sensitive to blood epinephrine

# $\alpha_2$ Receptors

- 1) Located on **presynaptic** axons.
- 2) When stimulated, result in **inhibition** of norepinephrine release in the synapse:
  - A. A **negative-feedback system**
  - B. Some drugs to **lower blood pressure** act on these  $\alpha_2$  receptors to inhibit presynaptic neurons in the brain, inhibiting the whole **sympathoadrenal** system.
  - C. There are different subtypes that will give **different responses**

# Response to Cholinergic Stimulation

- ACh released from preganglionic neurons of both the sympathetic and **parasympathetic division is stimulatory**.
- ACh from postganglionic neurons of the parasympathetic division is usually **stimulatory**, but some are **inhibitory**, depending on receptors.
- In general, sympathetic and parasympathetic effects are **opposite**

# Cholinergic Receptors

**A. Nicotinic:** found in autonomic ganglia

- 1) Stimulated by Ach from **preganglionic neurons**
- 2) Serve as ligand-gated ion **channels** for  $\text{Na}^+$  &  $\text{K}^+$

**B. Muscarinic:** found in visceral organs and stimulated by **release of Ach from postganglionic neurons**

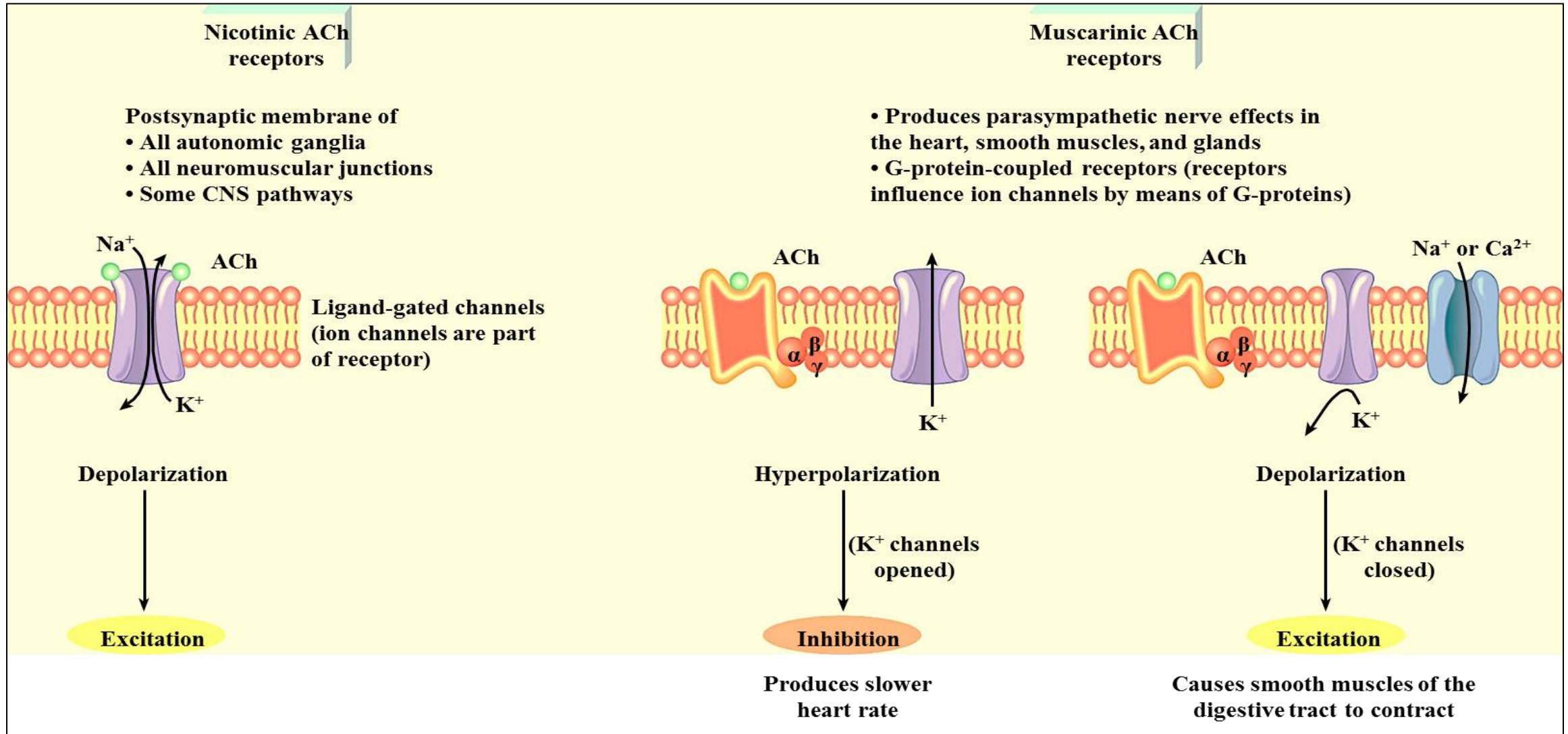
- 1) Five types identified; can be stimulatory or inhibitory (**opening  $\text{K}^+$  or  $\text{Ca}^{2+}$  channels**)
- 2) Use G-proteins and **second messenger system**



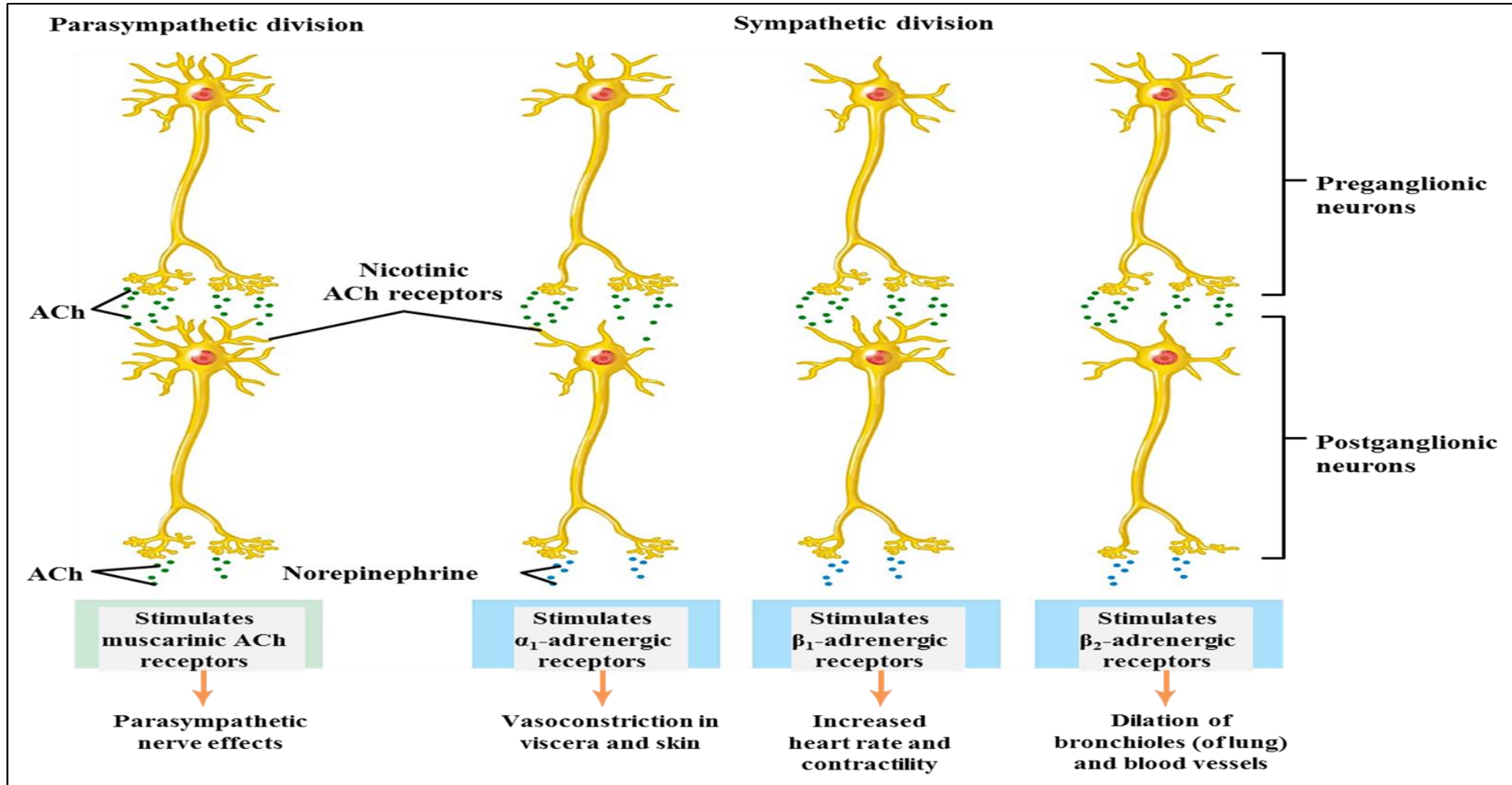
**Table 9.6 | Cholinergic Receptors and Responses to Acetylcholine**

Receptor	Tissue	Response	Mechanisms
Nicotinic	Skeletal muscle	Depolarization, producing action potentials and muscle contraction	ACh opens cation channel in receptor
Nicotinic	Autonomic ganglia	Depolarization, causing activation of postganglionic neurons	ACh opens cation channel in receptor
Muscarinic (M <sub>3</sub> , M <sub>5</sub> )	Smooth muscle, glands	Depolarization and contraction of smooth muscle, secretion of glands	ACh activates G-protein coupled receptor, opening Ca <sup>2+</sup> channels and increasing cytosolic Ca <sup>2+</sup>
Muscarinic (M <sub>2</sub> )	Heart	Hyperpolarization, slowing rate of spontaneous depolarization	ACh activates G-protein coupled receptor, opening channels for K <sup>+</sup>

# Comparison of Nicotinic & Muscarinic ACh Receptors



# Receptor Activity in Autonomic Regulation



# Organs with Dual Innervation

- Most visceral organs are innervated by both sympathetic and parasympathetic neurons.
1. Antagonistic (most common)
  2. Complementary
  3. Cooperative

# Antagonistic Effects

## **1. Heart rate:**

sympathetic increases, parasympathetic decreases.

## **2. Digestive functions:**

sympathetic decreases, parasympathetic increases.

## **3. Pupil diameter:**

sympathetic dilation, parasympathetic constriction.



- **Complementary Effects:** Occur when both divisions produce similar effects on the same target.
- Example: Salivary gland secretion: Parasympathetic division stimulates secretion of watery saliva; sympathetic constricts blood vessels so the secretion is thicker.
- **Cooperative Effects:** Occur when both divisions produce different effects that work together to promote a single action.
- Example – Effect of sympathetic and parasympathetic systems on the male and female reproductive systems.

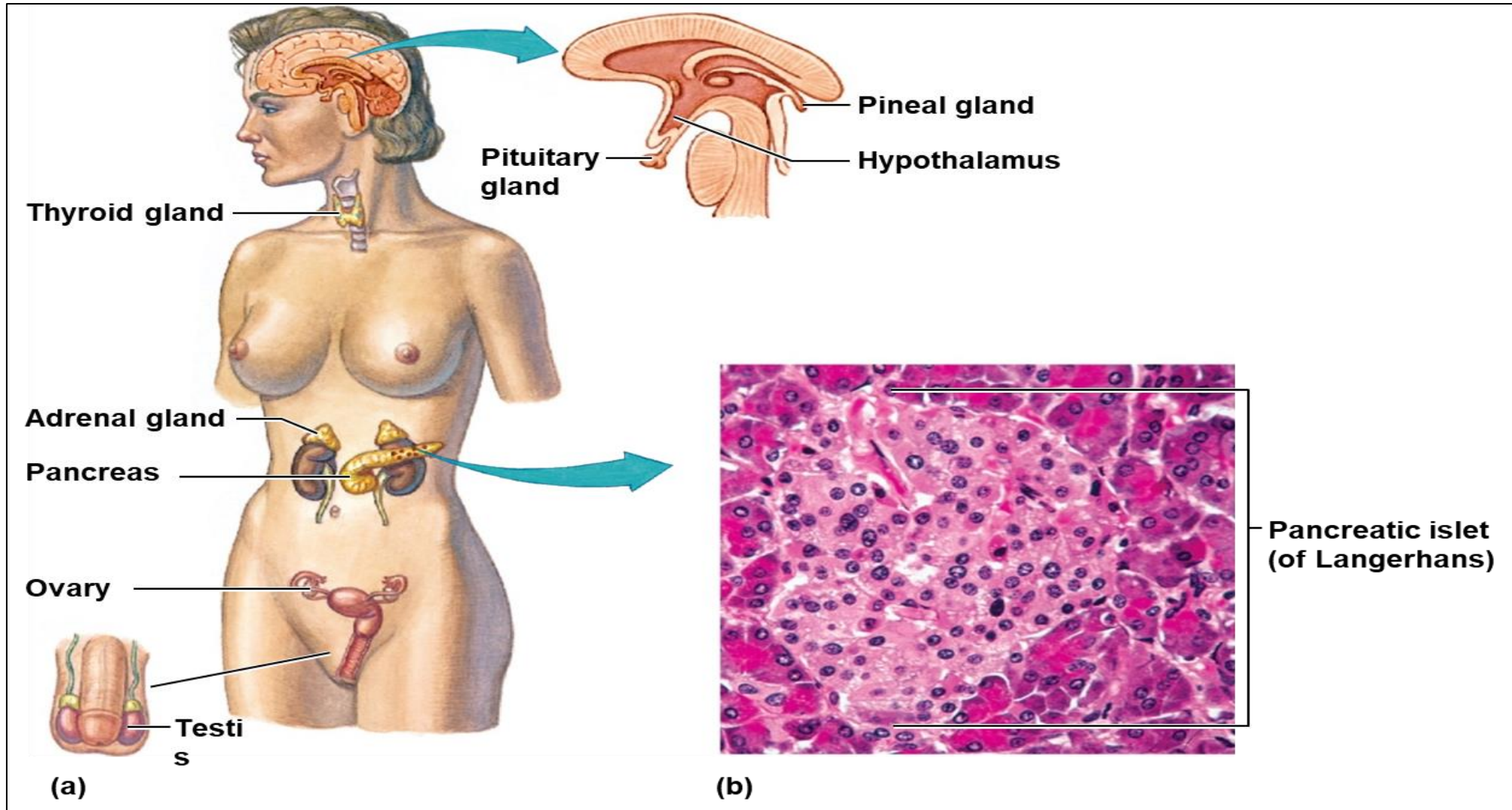
# Organs without Dual Innervation

- The following organs are innervated by the **sympathetic division only**:
  - A. Adrenal medulla
  - B. Arrector pili muscles in skin
  - C. Sweat glands in skin
  - D. Most blood vessels
- Regulated **by increase and decrease in sympathetic nerve activity**.
- Important for body temperature **regulation through blood vessels and sweat glands**

# Chapter 11: Endocrine Glands and Hormones

1. Ductless
2. Secrete hormones into the blood
3. Hormones are carried to target cells having receptors for those hormones.
4. Neurohormones are secreted by specialized cells of the hypothalamus
5. Hormones help regulate body metabolism, growth, and reproduction

# Major Endocrine Glands



# Chemical Classification of Hormones

1. **Amines**, derived from tyrosine and tryptophan
  - ❖ Examples: hormones from the adrenal medulla, thyroid, and pineal glands.
2. **Polypeptides and proteins** :A peptide is two or more amino acids joined together by peptide bonds, and a polypeptide is a chain of many amino acids. A protein contains one or more polypeptides. Therefore, proteins are long chains of amino acids held together by peptide bonds.
  - ❖ Examples: antidiuretic hormone, insulin, and growth hormone
3. **Glycoproteins are long polypeptides bound to a carbohydrate.**
  - ❖ Examples: follicle-stimulating and luteinizing hormones (FSH, LH), Thyroid-stimulating hormone (TSH).
4. **Steroids are lipids derived from cholesterol**
  - ❖ Examples: testosterone, estradiol, progesterone, cortisol
  - ❖ Secreted by adrenal cortex and gonads



## Hormone Classifications by action

1. **Polar hormones: water soluble**
  - a. Cannot pass through plasma membranes.
  - b. Includes polypeptides, glycoproteins, catecholamines (norepinephrine and epinephrine).
  
2. **Nonpolar: insoluble in water**
  1. Often called lipophilic hormones.
  2. Can enter target cells directly.
  3. Include steroids, thyroid hormone, and melatonin.

# Hormone Interactions

**1. Synergistic Effects:** Occur when two or more hormones work together to produce a particular effect

A. **Effects may be additive**, each hormone act to produce same effect; as when epinephrine and norepinephrine each affect the heart **in the same way (increase heart rate)**.

B. **Effects may be complementary**, as when each hormone contributes a different piece of an overall outcome. (**Each** hormone mediates **significant function** while **all** hormones act together to perform **new complementary function**).

**For example**, producing milk requires estrogen, cortisol, prolactin, and oxytocin.

**2. Permissive Effects:** Occur when one hormone makes the target cell more responsive (enhances responsiveness of the target cell) to a second hormone.

A. Exposure to estradiol makes the uterus more responsive to progesterone.

B. Increased secretion of PTH makes the intestines more responsive to Vitamin D<sub>3</sub> in calcium absorption. This is achieved when PTH helps in activating Vitamin D<sub>3</sub> (Prehormone- Hormone).

**e.g. Parathyroid hormone (PTH)** stimulates the **production** hydroxylating enzymes that activate **vitamin D3**.

## **3. Antagonistic Effects**

A. Occur when hormones work in opposite directions.

B. Insulin and glucagon both affect adipose tissue.

- Insulin stimulates fat storage

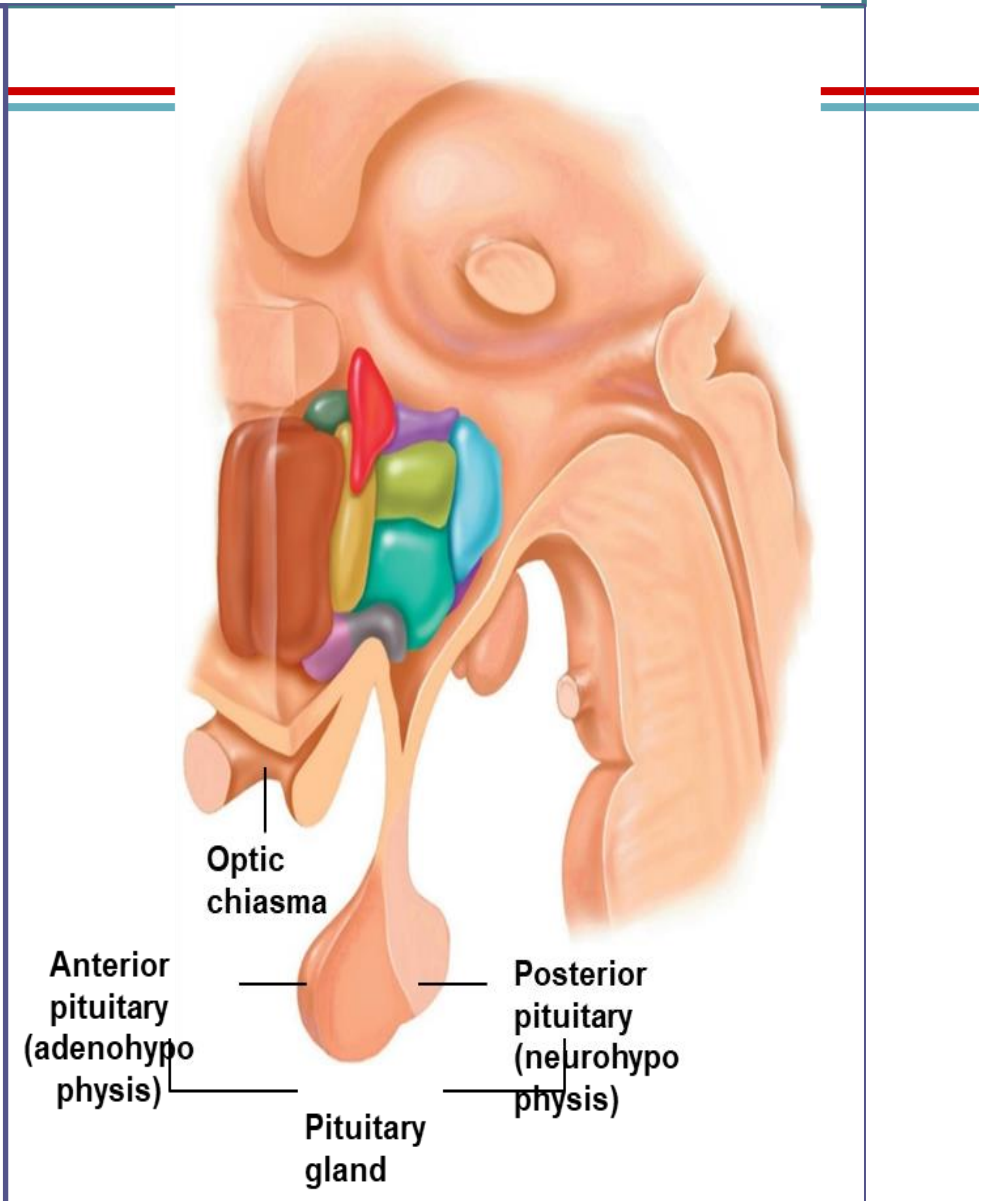
- Glucagon stimulates fat breakdown.

# Mechanisms of Hormone Action: Introduction

1. **Hormones bind** to receptors **on** or **in** target cells.
  - a. Binding is **highly specific**.
  - b. Hormones bind to receptors with **a high affinity**.
  - c. Hormones bind to receptors with a **low capacity**; saturating the receptors with hormone molecules
2. **Lipophilic hormone** (non-polar) receptors are in the **cytoplasm** or **nucleus**
3. **Water-soluble** (polar) hormone receptors are on **the outer surface** of the plasma membrane

# Pituitary gland

1. The pituitary gland is **attached to the hypothalamus by the infundibulum**.
2. Divided into an **anterior lobe (adenohypophysis)** and a **posterior lobe (neurohypophysis)**.
  - a. The anterior pituitary **is glandular epithelium**
  - b. The posterior pituitary **is nervous tissue**



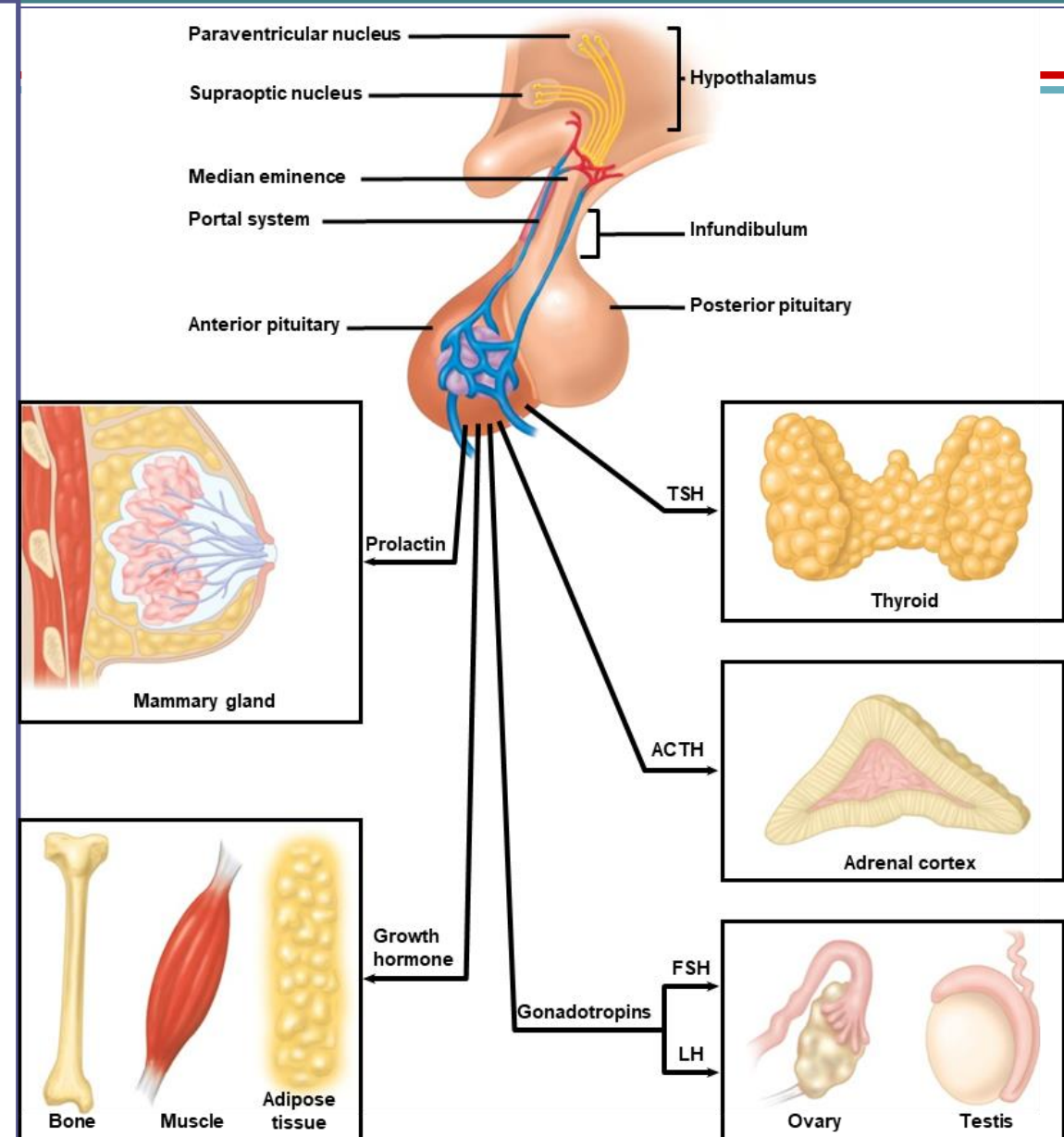
## Posterior Pituitary Hormones

- **Stores and releases two hormones made in the hypothalamus:**
  - 1) **Antidiuretic hormone (ADH)**, which promotes the retention of water in the kidneys (also called arginine vasopressin – AVP)
  - 2) **Oxytocin**, which stimulates contractions in childbirth and milk let-down in lactation.
- **Hypothalamic Control of the Posterior Pituitary**
  - 1) **ADH and oxytocin** are produced **by the supraoptic and paraventricular nuclei** of the **hypothalamus**, respectively
  - 2) They are transported along axons of the **hypothalamo-hypophyseal tract** to the **posterior pituitary** where they are **stored**.



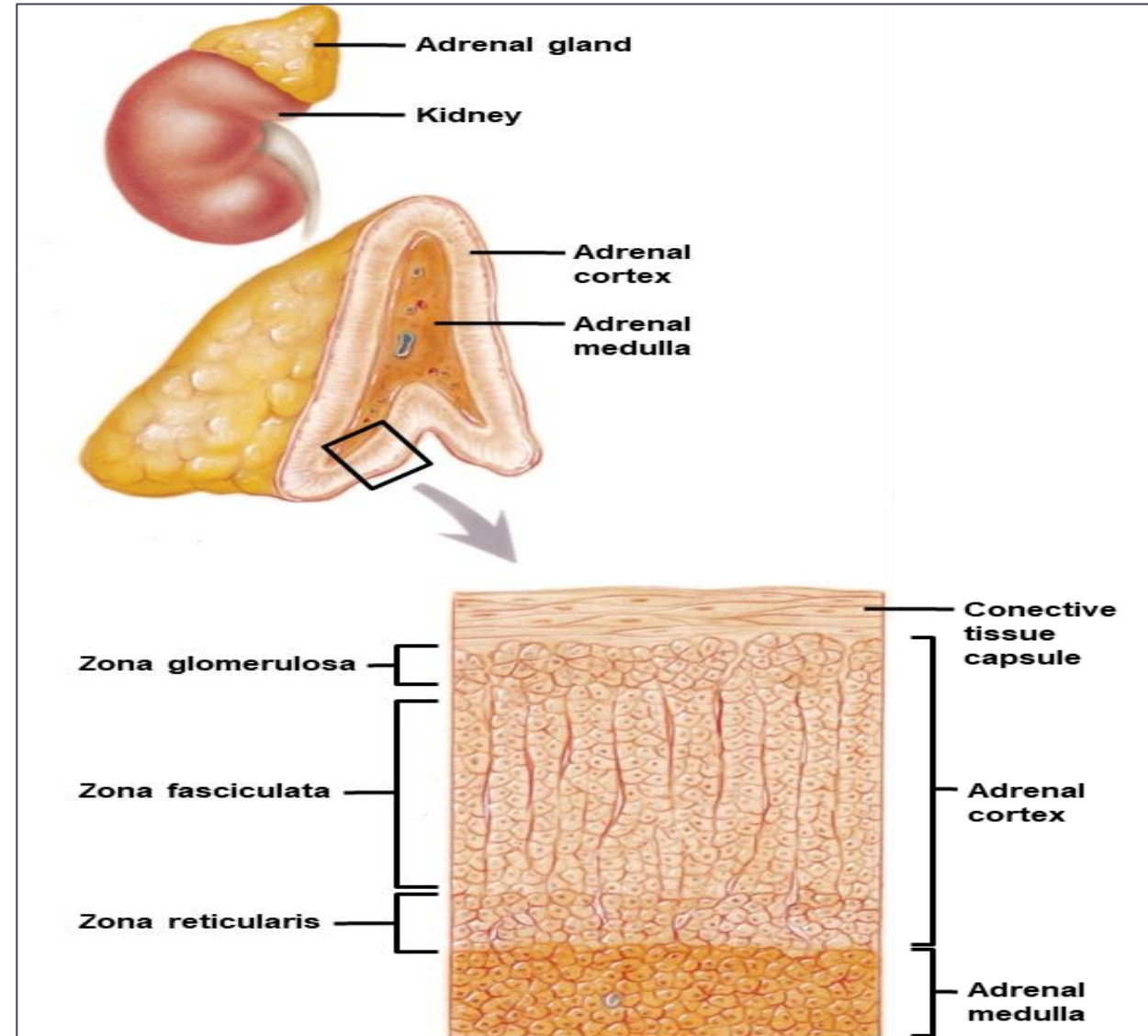
# Anterior Pituitary Hormones

- Secreted by the **anterior lobe**
- **Trophic hormones** stimulate hormone secretion in other glands:
  - a. **Growth hormone (GH)** or **somatotropin**
  - b. **Thyroid-stimulating hormone (TSH)** or **thyrotropin**
  - c. **Adrenocorticotrophic hormone (ACTH)** or **corticotropin**
  - d. **Follicle-stimulating hormone (FSH)** or **folliculotropin**
  - e. **Luteinizing hormone (LH)** or **luteotropin** – in the male, it is interstitial cell stimulating hormone (ICSH)
  - f. **Prolactin (PRL)**



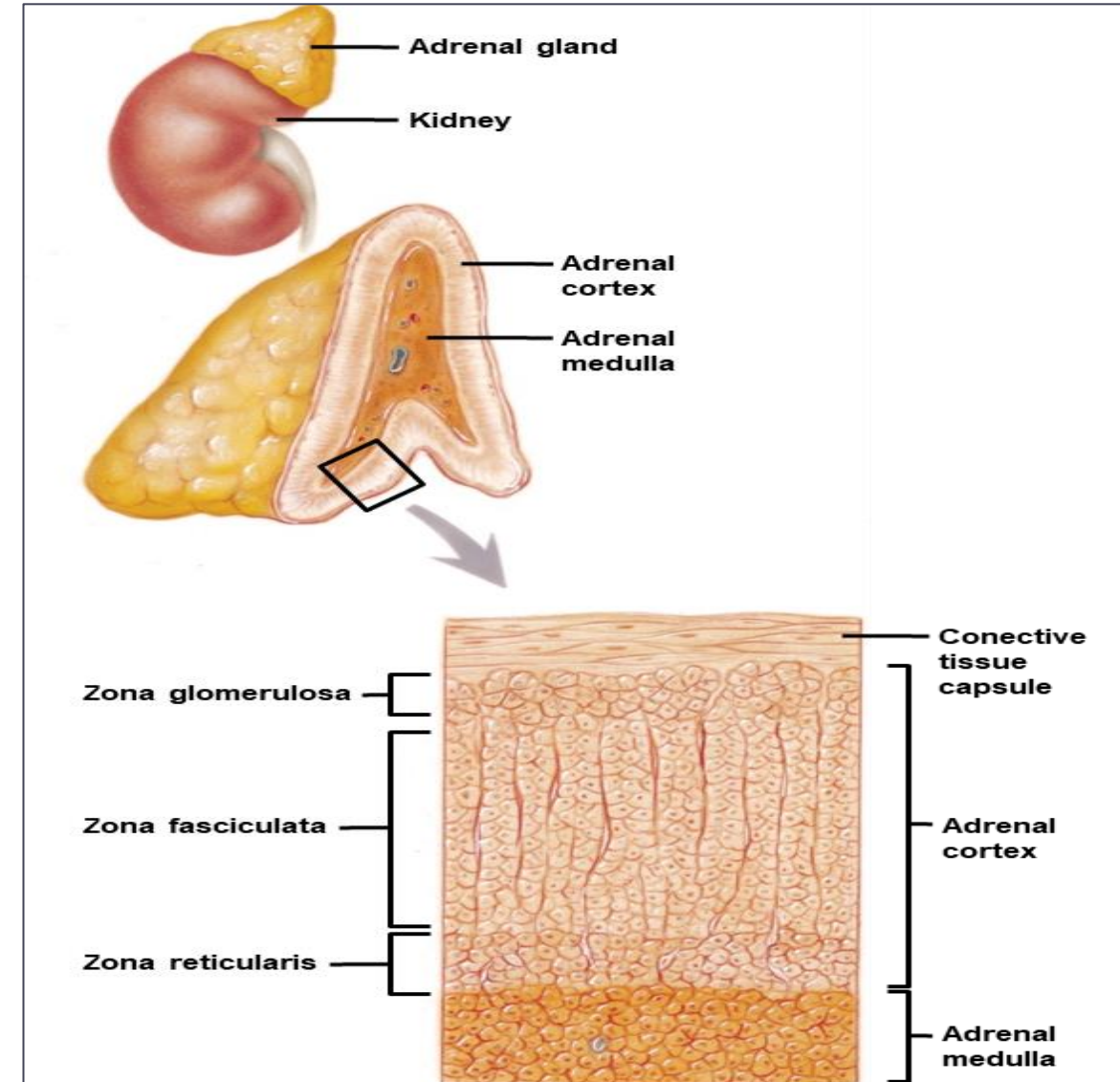
# Structure of the Adrenal Glands

1. Found atop the kidneys.
2. Consist of an **outer adrenal cortex** and an **inner adrenal medulla** that **function as separate glands**
  - a. The **adrenal medulla**
  - b. The **adrenal cortex**



# Structure of the Adrenal Glands

- A. The **adrenal medulla** is **neural tissue** and secretes **epinephrine** and **norepinephrine** in response to **sympathetic neural stimulation**.
- B. The **adrenal cortex** is **glandular epithelium** and secretes **steroid hormones** in response to **ACTH**; three layers – **zona glomerulosa**, **zona fasciculata**, and **zona reticularis**.



# Functions of the adrenal cortex

1. Secretes hormones made from **cholesterol**; called **corticosteroids** or **corticoids**
2. **Three categories:**
  - A. **Mineralocorticoids** from the zona glomerulosa regulate  $\text{Na}^+$  and  $\text{K}^+$  balance. Example: **aldosterone**.
  - B. **Glucocorticoids** from the zona fasciculata regulate glucose metabolism. Example: **cortisol**.
  - C. **Adrenal androgens** from the zona reticularis are weak sex hormones that supplement those made in the gonads. **Ex - DHEA**



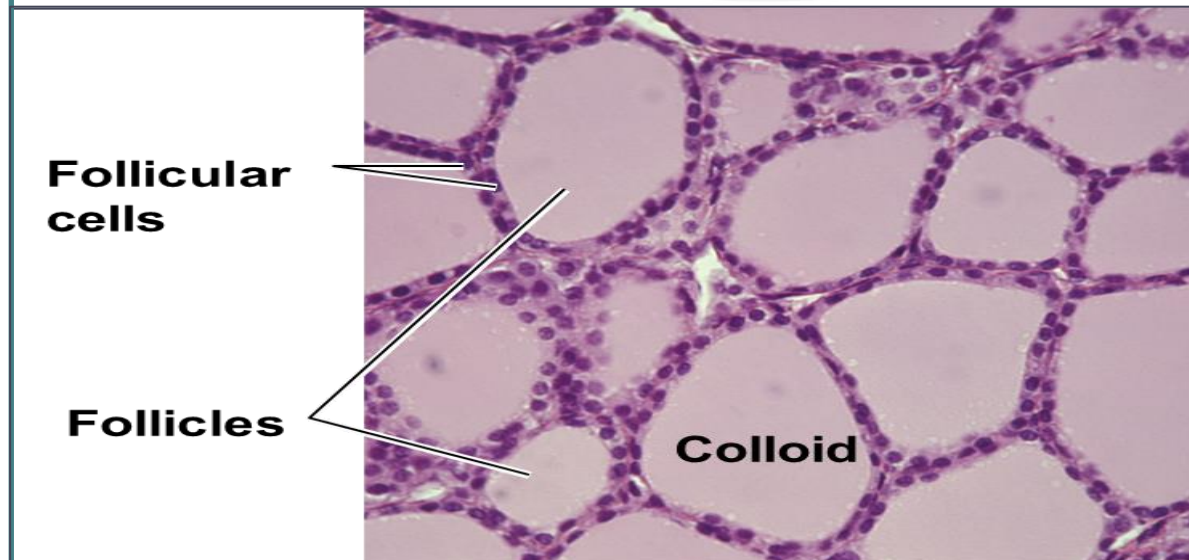
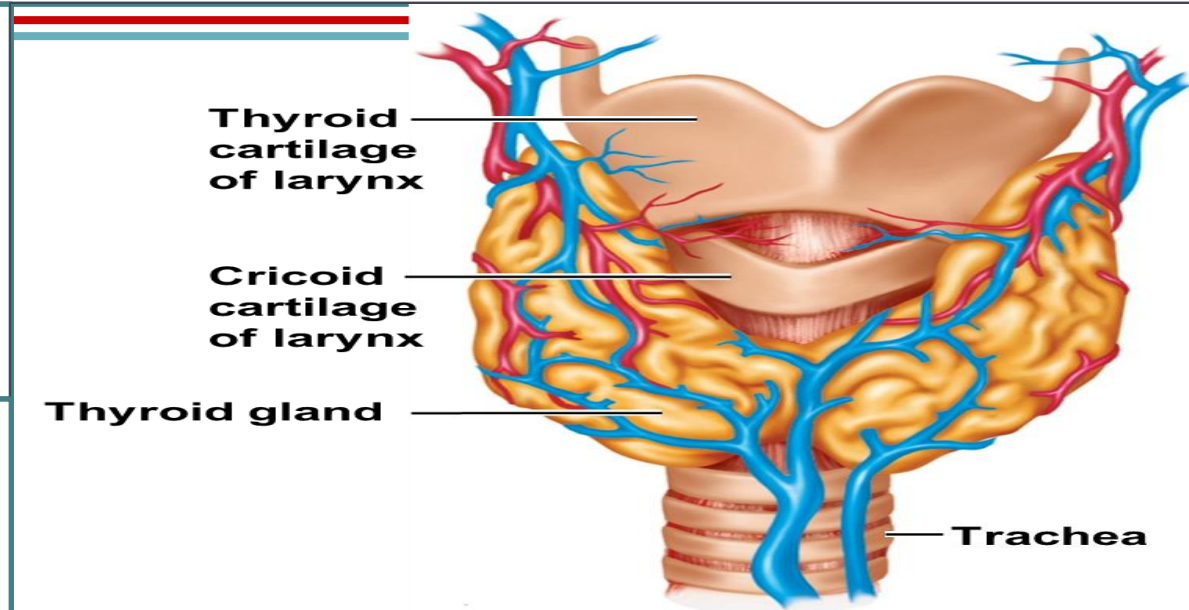
# Thyroid Gland Structure

- **Thyroid Gland Structure:**
  1. Located just below the larynx
  2. Has two lobes on either side of the trachea, connected by the isthmus

- **Microscopic Thyroid Gland Structure:**

Consists of hollow spaces called thyroid follicles lined with **simple cuboidal epithelium** composed of **follicular cells** that produce **thyroxine**.

1. Interior of the follicles is filled with a fluid called **colloid**.
2. Outside of the follicles are **parafollicular cells** that secrete **calcitonin**





## Production of Thyroid Hormone

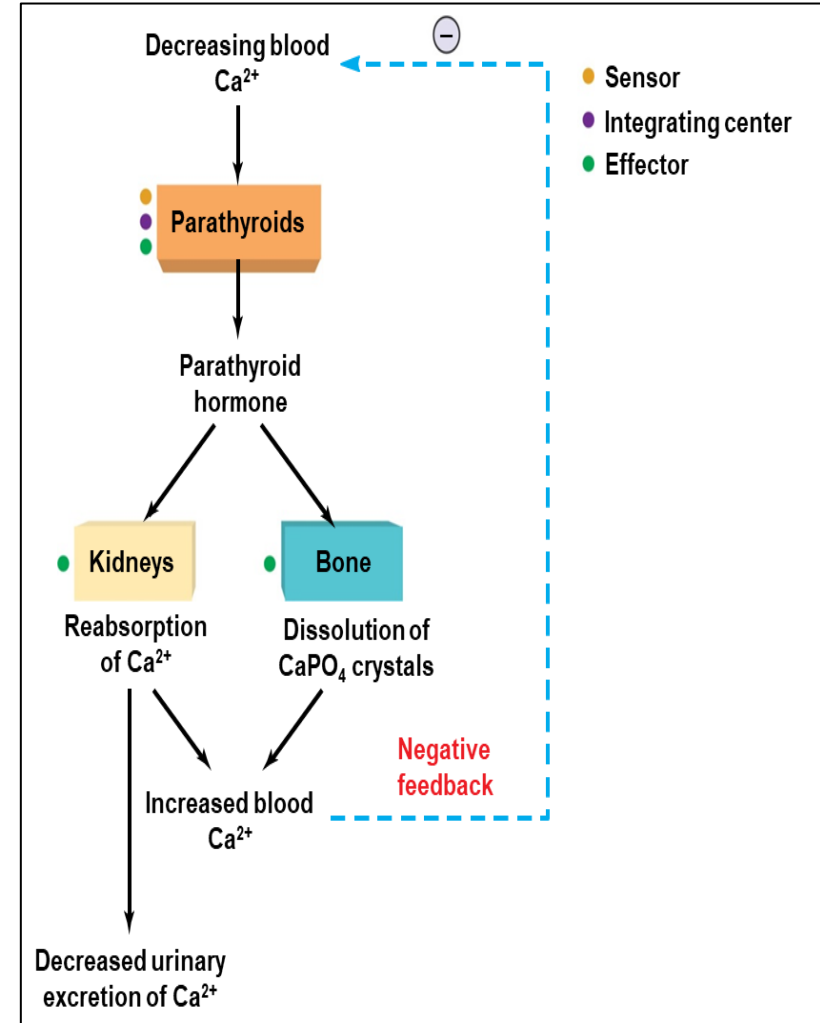
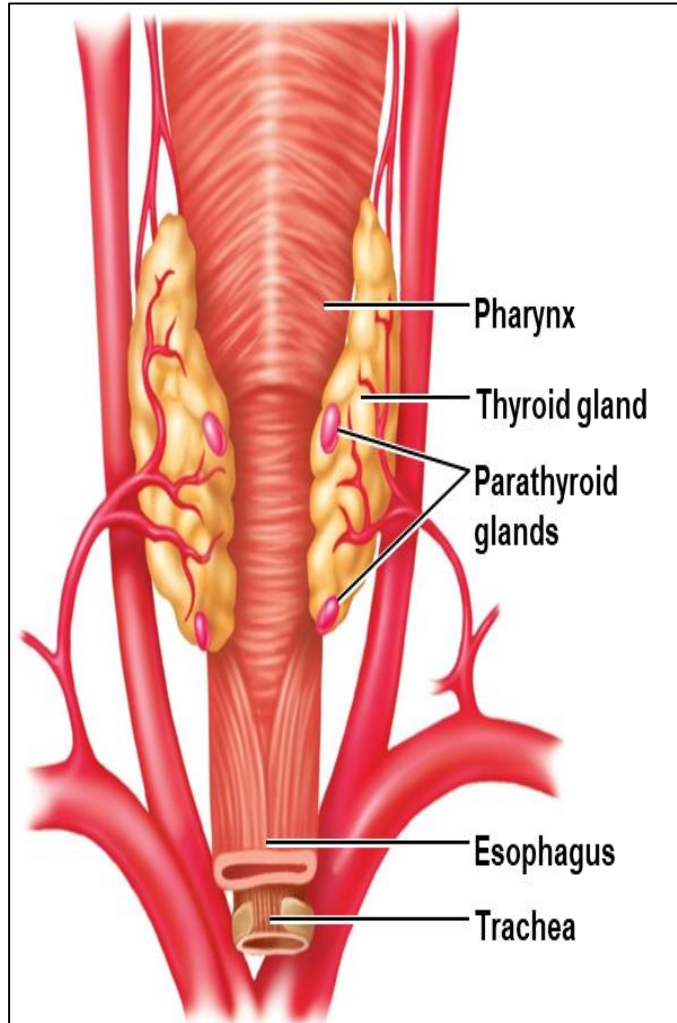
- $T_4$  (tetraiodothyronine or thyroxine)
- $T_3$  (triiodothyronine)
- They are secreted into the blood when the thyroid gland is stimulated by TSH.
- Action of Thyroid Hormone:
  - A. Stimulates protein synthesis
  - B. Promotes maturation of the nervous system
  - C. Increases rates of cellular respiration
  - D. Elevates basal metabolic rate

# Calcitonin

1. Also called **thyrocalcitonin**.
2. Made by the **parafollicular cells**.
3. Inhibits **dissolution of calcium from bone** and stimulates excretion of **calcium in the kidneys to lower blood calcium levels**.

# Parathyroid Glands

- Generally, 4 glands embedded in the back of the thyroid gland.
- Secrete **parathyroid hormone (PTH)**.
- Hormone **promotes a rise in blood calcium** by acting on **bones, kidneys, and intestine**.



# **Chapter 10:**

# **Sensory physiology**

# Characteristics of Sensory Receptors

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1. **Sensory receptors transduce** (change) different forms of energy in the “real world” into nerve impulses.
2. **Different modalities of** sensations (sound, light, pressure) arise from differences in neural pathways and synaptic connections



# Categories of sensory receptors

- Functional categories (according to the type of signal they transduce):
  - 1) **Chemoreceptors**: sense chemicals in the environment (taste, smell) or blood.
  - 2) **Photoreceptors**: sense light.
  - 3) **Thermoreceptors**: respond to cold or heat.
  - 4) **Mechanoreceptors**: stimulated by mechanical deformation of the receptor (touch, hearing).
  - 5) **Nociceptors**: Pain receptors that depolarize when tissues are damaged (stimuli can include heat, cold, pressure, or chemicals).

# Sensory Receptor Categories

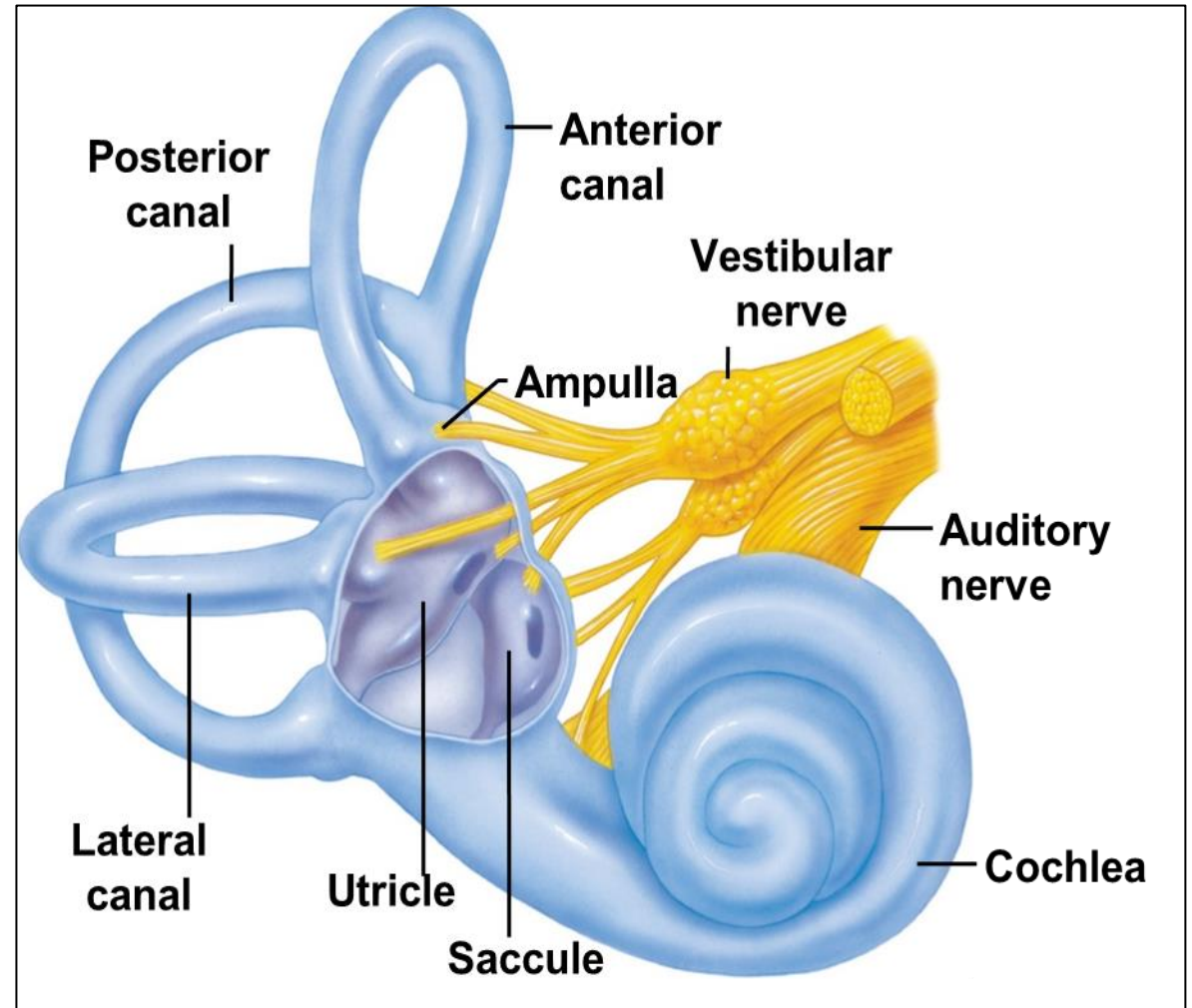
- **Receptors can be classified** by the type of information they deliver to the brain:
  - 1) **Proprioceptors**: found in muscles, tendons, and joints. Provide a sense of body position and allows fine muscle control.
  - 2) **Cutaneous (skin) receptors** – touch, pressure, heat, cold, and pain.
  - 3) **Special senses** – vision, hearing, taste, smell, equilibrium

# Sensory Receptor Categories

- Receptors can be classified by the origin of the information:
  - 1) **Exteroceptors**: respond to stimuli from outside the body; includes cutaneous receptors and special senses.
  - 2) **Interoceptors**: respond to internal stimuli; found in organs; monitor blood pressure, pH, and oxygen concentrations.

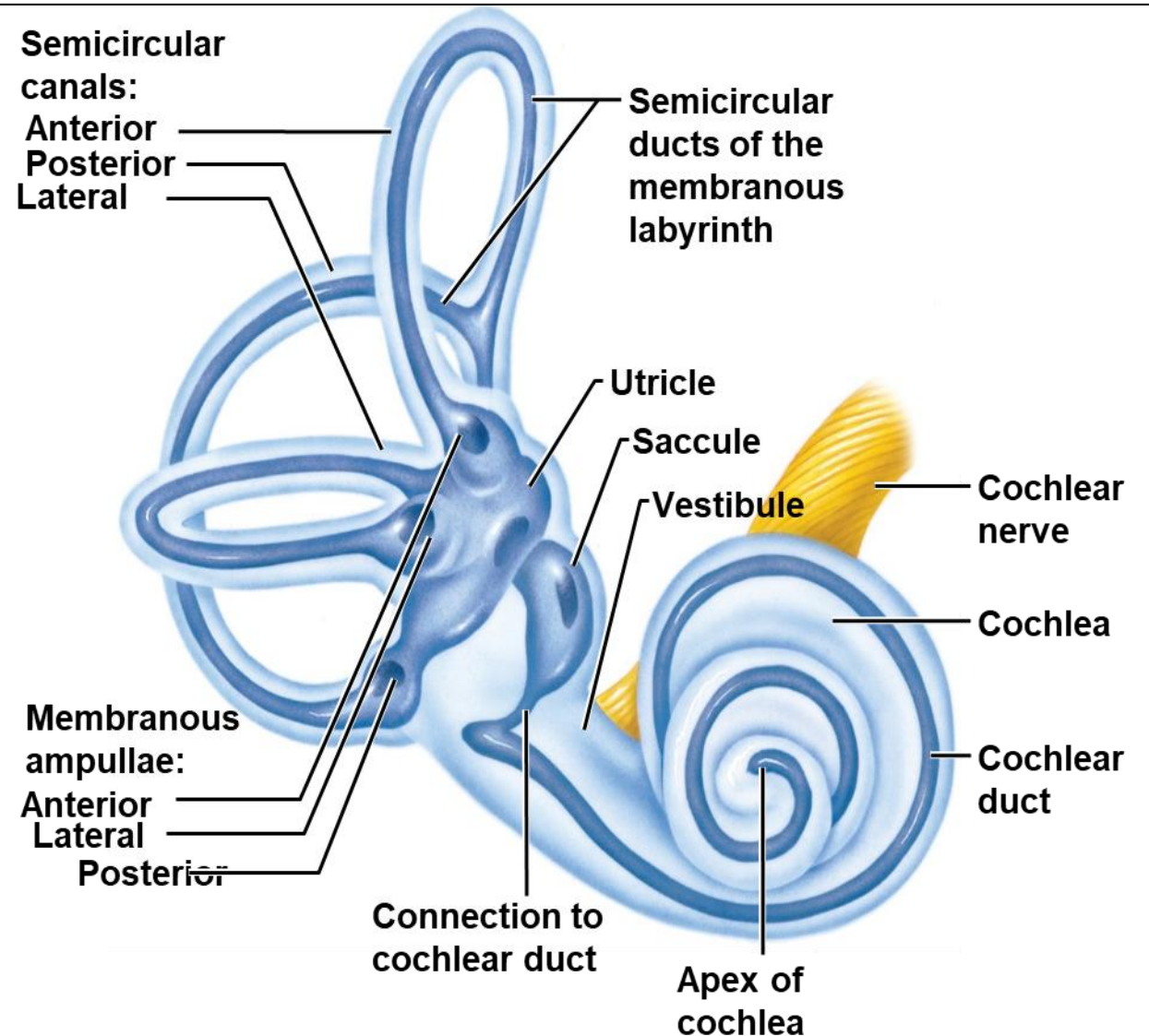
# Vestibular Apparatus and Equilibrium

1. **Provides** a sense of equilibrium
2. **Located** in the inner ear
3. Consists of:
  - a. **Otolith organs**  
Utricle and saccule – linear acceleration
  - b. **Semicircular canals** – rotational acceleration



# Labyrinths of the Inner Ear

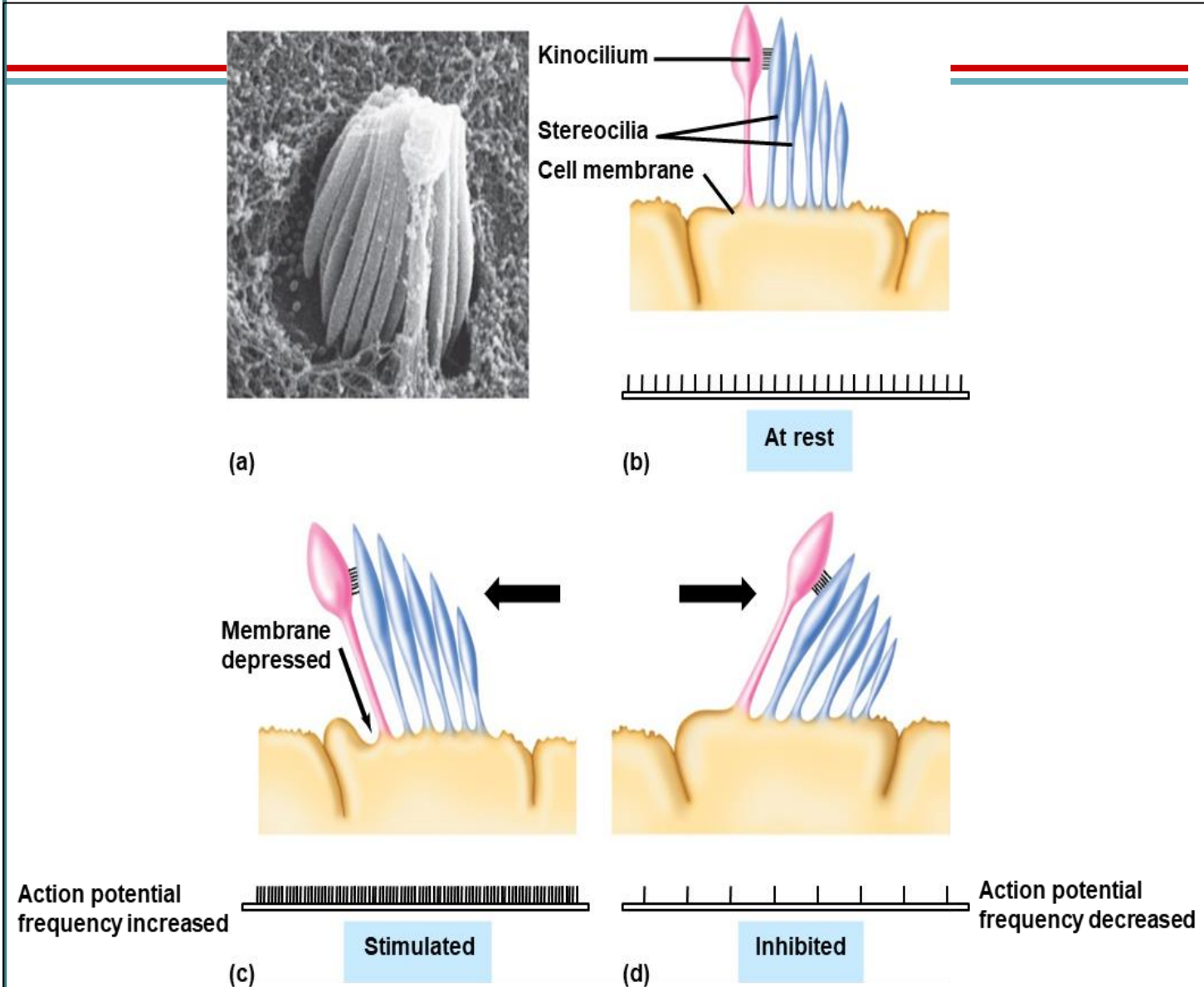
- A. **Consists of** a bony labyrinth surrounding a membranous labyrinth.
- B. **Between the** two is fluid called perilymph.
- C. **Within the membranous** labyrinth is fluid called endolymph.
- D. **Endolymph has** an unusually high  $K^+$  concentration, which will produce depolarization





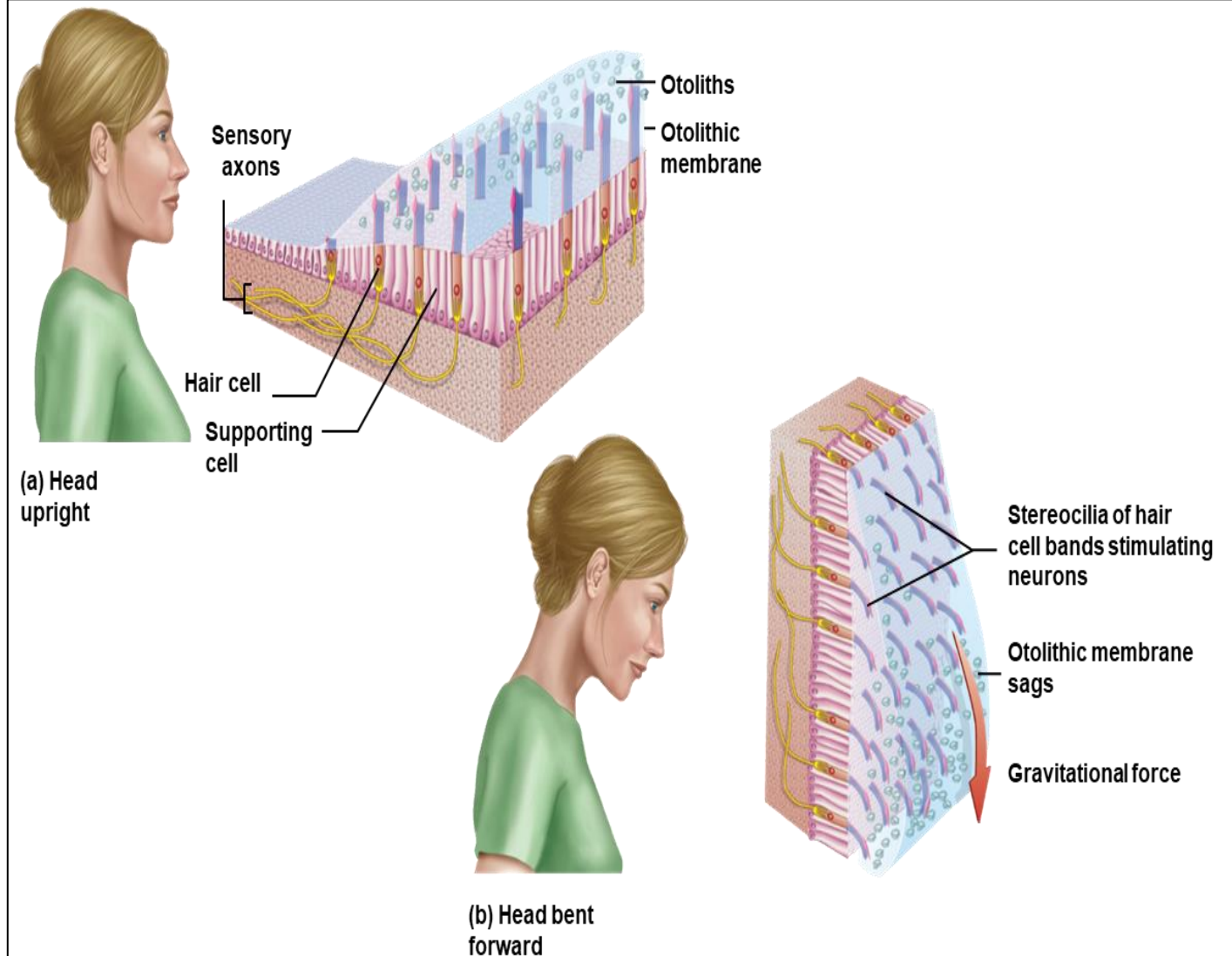
# Sensory Hair Cells

1. **Modified epithelial cells with 20–50** hairlike extensions called stereocilia (not true cilia) and one kinocilium (true cilium).
2. **When stereocilia** bend toward the kinocilium,  $K^+$  channels open, and  $K^+$  rushes into the cell and depolarizes the cell.
3. **Cells release a neurotransmitter** that depolarizes sensory dendrites in the vestibulocochlear nerve.
4. **Bending away from the kinocilium** hyperpolarizes sensory dendrites.
5. **Code for detection of direction.**



# Otolithic Organs: Utricle and Sacculle

1. **Provide information** about linear acceleration:
  - a. **Utricle**: horizontal
  - b. **Sacculle**: vertical
2. **Specialized epithelium** called the macula houses hair cells.
  - a. **Stereocilia** are embedded in a gelatinous otolithic membrane.
  - b. **The gel also** contains crystals of calcium carbonate called otoliths (ear stones)



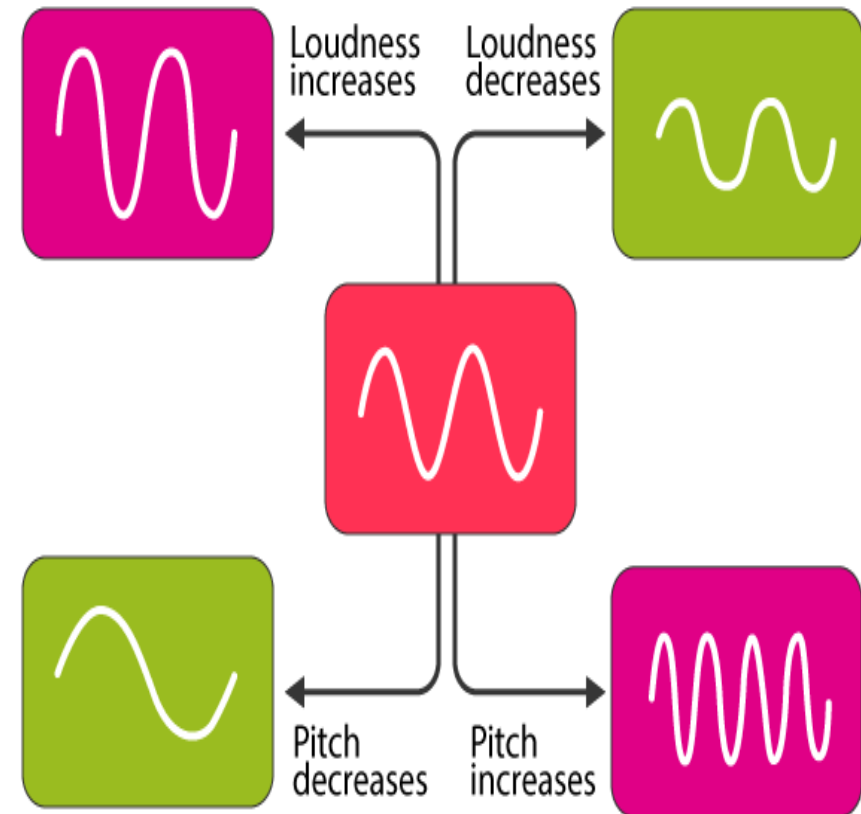
# Semicircular Canals

1. **Project along three** planes to detect rotation:
  - a. **Each canal** contains a semicircular duct filled with endolymph.
  - b. **At the base** of each duct is an enlarged area called the ampulla.

# The Ears and Hearing: Sound Waves

- **Characterized by:**
  - A. **Frequency**, measured in hertz (Hz). Higher frequencies have higher pitches.
    - 1) **Human range is 20–20,000 Hz.**
  - B. **Intensity or loudness**, measured in decibels
    - 1) **Related** to the amplitude of the wave
    - 2) **Human optimal range is 0-80 dB**

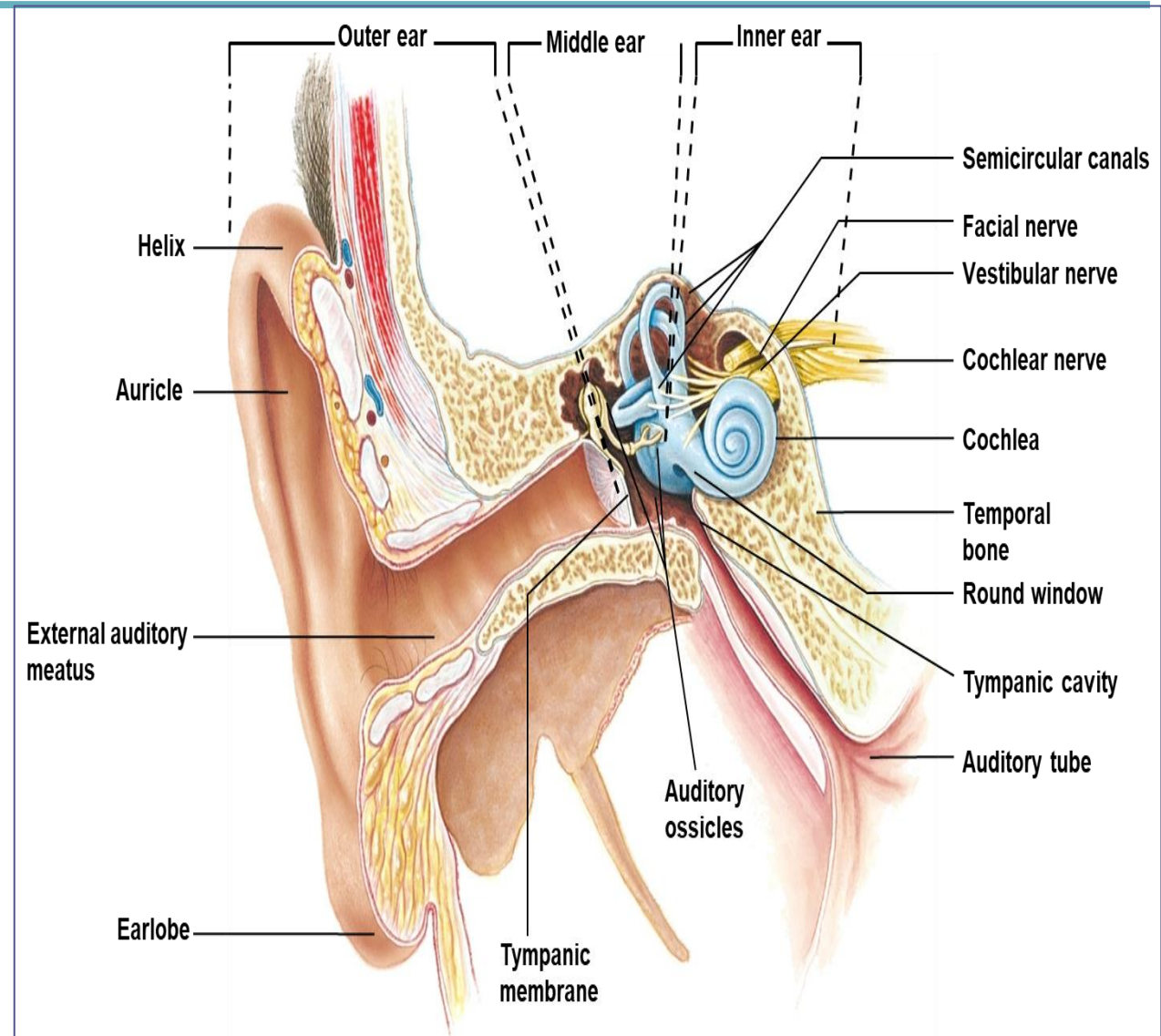
## DIFFERENCE BETWEEN PITCH AND LOUDNESS



# The Ear

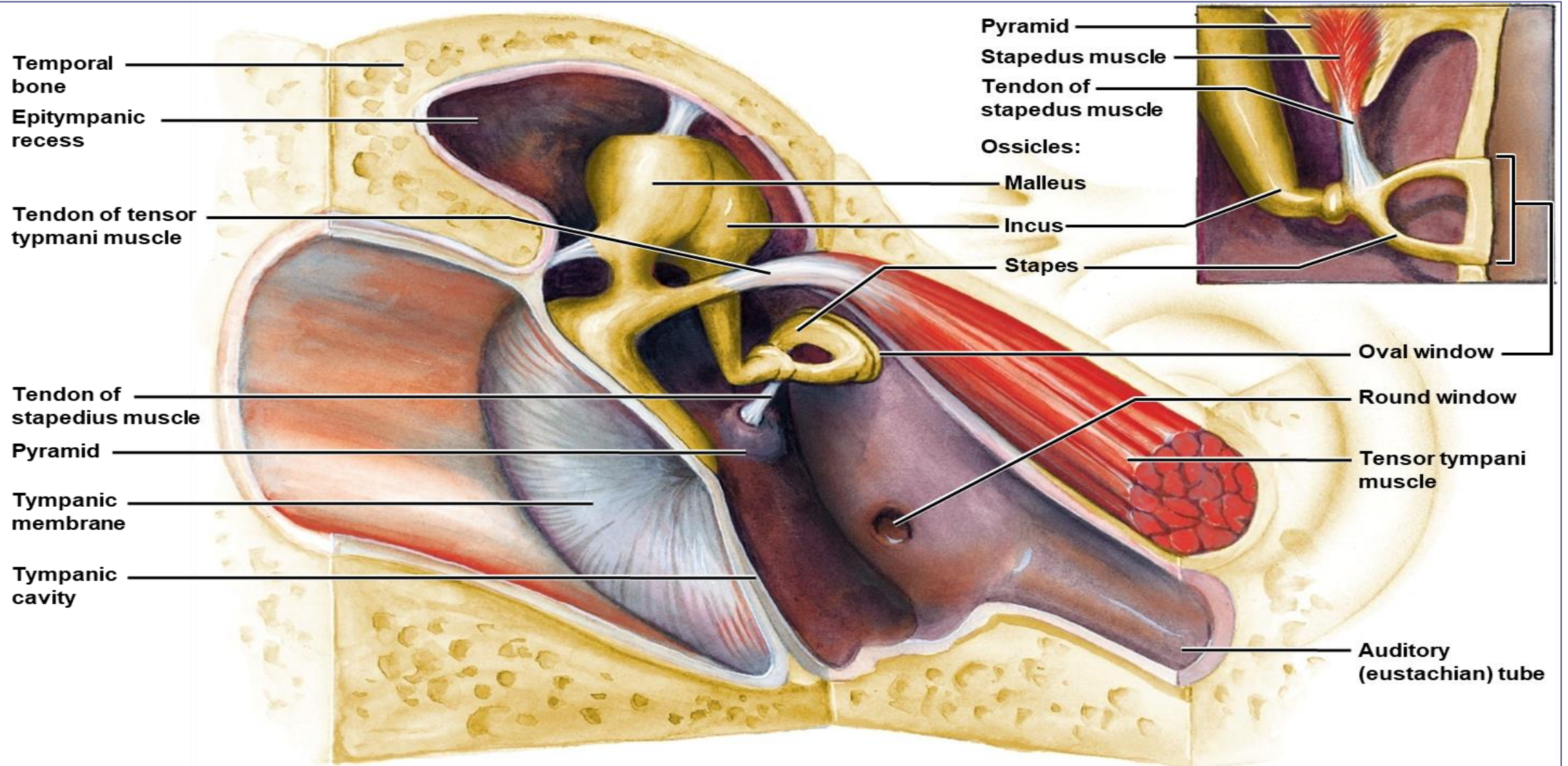
## Outer Ear and Middle Ear

1. Sound waves are funneled by the pinna (or auricle) into the external auditory meatus, which channels them to the tympanic membrane (eardrum).
2. Air-filled cavity between the tympanic membrane and the cochlea.
3. Contains three bones called ossicles:
  - a. Malleus incus stapes
  - b. Vibrations are transmitted and amplified along the bones.
  - c. The stapes is attached to the oval window, which transfers the vibrations into the cochlea.
  - d. Stapedius muscle dampens the stapes if the sound is too intense





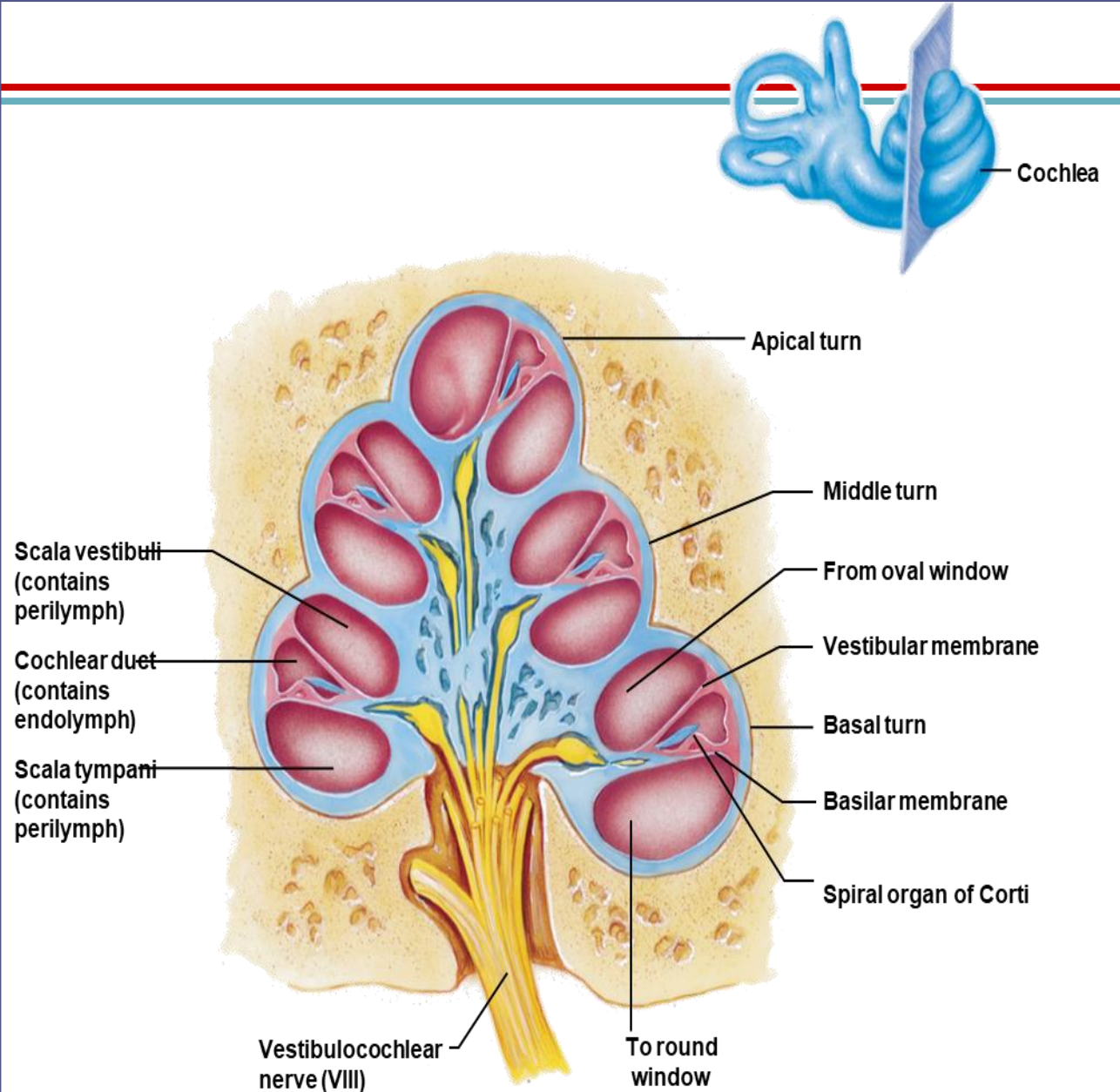
# Medial View of the Middle Ear





# Inner Ear: the Cochlea is responsible for hearing

- Three chambers:
  - a. The upper chamber is a portion of the bony labyrinth called the **scala vestibuli**.
  - b. There is also a lower bony chamber called the **scala tympani**.
  - c. Both chambers are filled with **perilymph**.
  - d. The cochlea also contains a portion of the membranous labyrinth called the **scala media, or cochlear duct, filled with endolymph**.
- ❖ **Middle chamber** that coils in three turns.



# Sound Transmission

- A. **Vibrations from the oval** window of the middle ear displace perilymph in the scala vestibuli.
- B. **Vibrations pass through** the vestibular membrane into the cochlear duct through the endolymph.
- C. **Next, vibrations pass** through the basilar membrane into the perilymph of the scala tympani.
- D. **Vibrations leave** the inner ear via the round window.
- E. **Sound waves are** transmitted through the cochlear duct at locations that depend on the frequency of the sound.
  - 1) Low-frequency sounds travel further down the spiral of the cochlea to the apex
  - 2) High-frequency sounds are closer the base

## Spiral Organ (Organ of Corti)

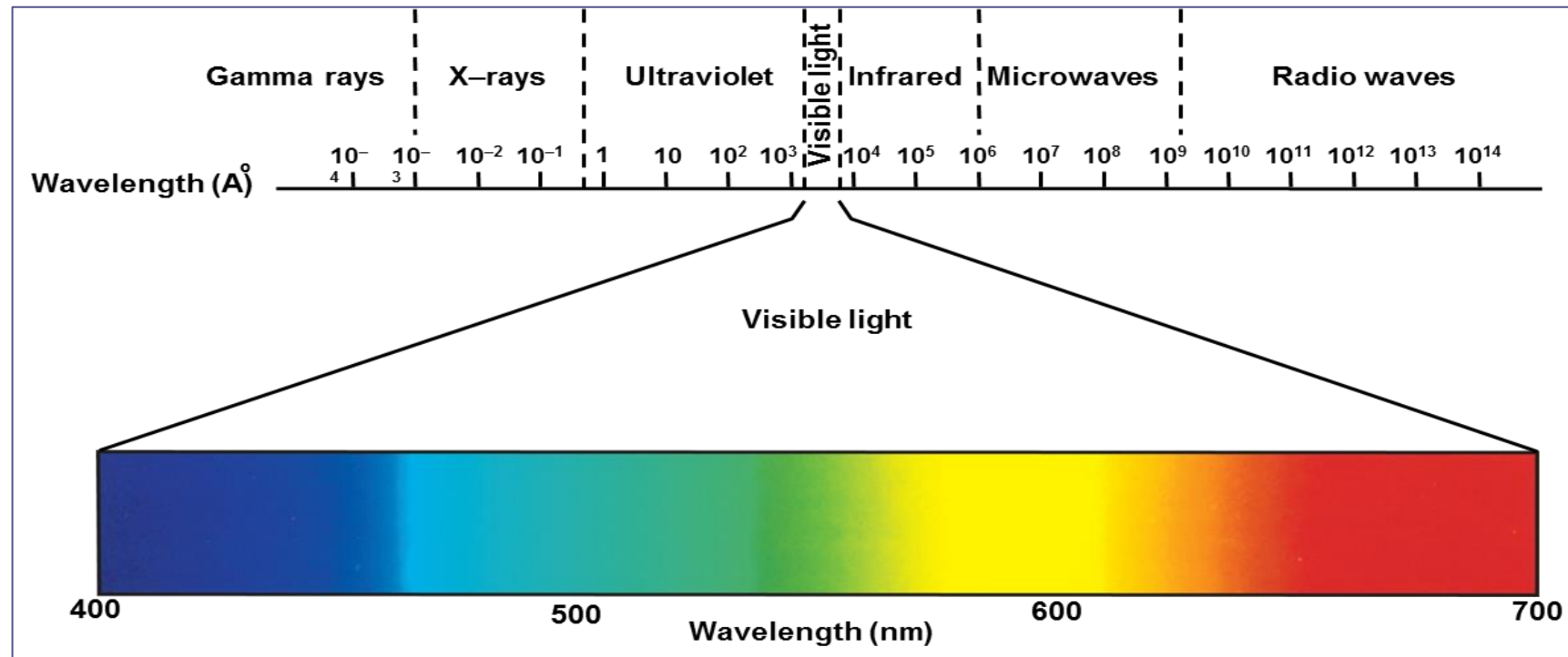
- **Sensory hair cells** are located on the basilar membrane, projecting into the endolymph of the cochlear duct
  - a. **Inner hair cells**: 3500 that form one row that runs the length of the basilar membrane. Each is innervated by 10–20 sensory neurons of cranial nerve VIII and relay sound.
  - b. **Outer hair cells**: 11,500 arranged in rows with 3 rows per turn. They are innervated by motor neurons that make them shorten when depolarized and elongate when hyperpolarized

# How hearing works

- a. When sound waves enter the scala media, the tectorial membrane vibrates, bending stereocilia.
  - 1) Opens  $K^+$  channels that are facing the endolymph
  - 2)  $K^+$  rushes in, depolarizing the cell
  - 3) Releases glutamate onto sensory neurons
  - 4)  $K^+$  returns to perilymph at the base of the stereocilia
- b. The greater the amount of basilar membrane displacement and bending the stereocilia, the more glutamate is released, producing a greater receptor potential.

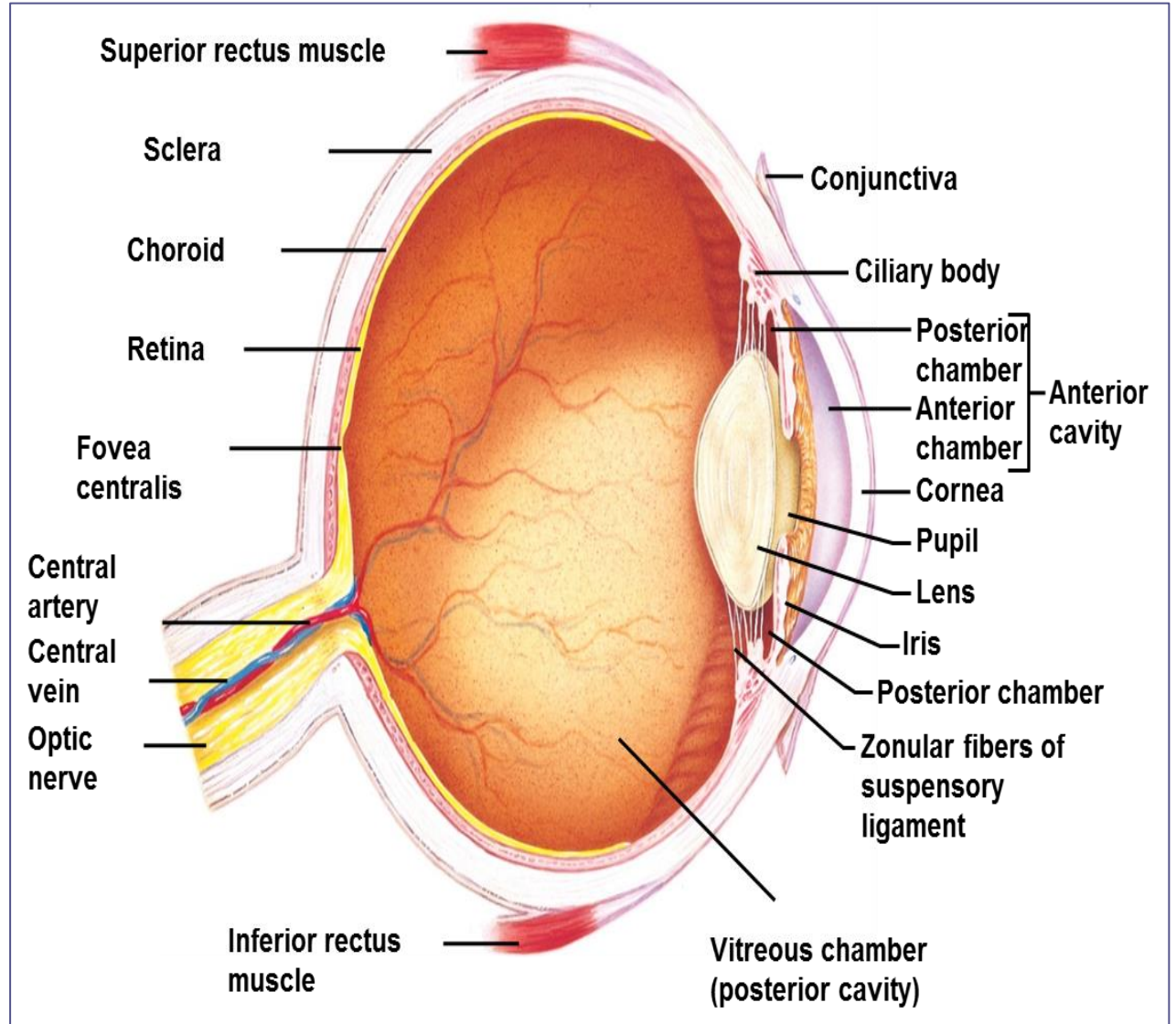
# The Eyes and Vision

1. Vision comes from light energy transduced into nerve impulses
2. Only a limited part of the electromagnetic spectrum can excite photoreceptors.



# Internal Anatomy of the Eyeball and general pathway of light through the eye

- a. Light passes through the cornea and into the anterior chamber of the eye.
- b. Next, it passes through the pupil, which can change shape (due to the pigmented iris muscle) to allow more or less light in.
- c. Then it passes through the lens, which can change shape to focus the image.
- d. Then passes through the posterior chamber and the vitreous body.
- e. Finally, it hits the retina, where photoreceptors are found and then absorbed by the pigmented choroid layer.



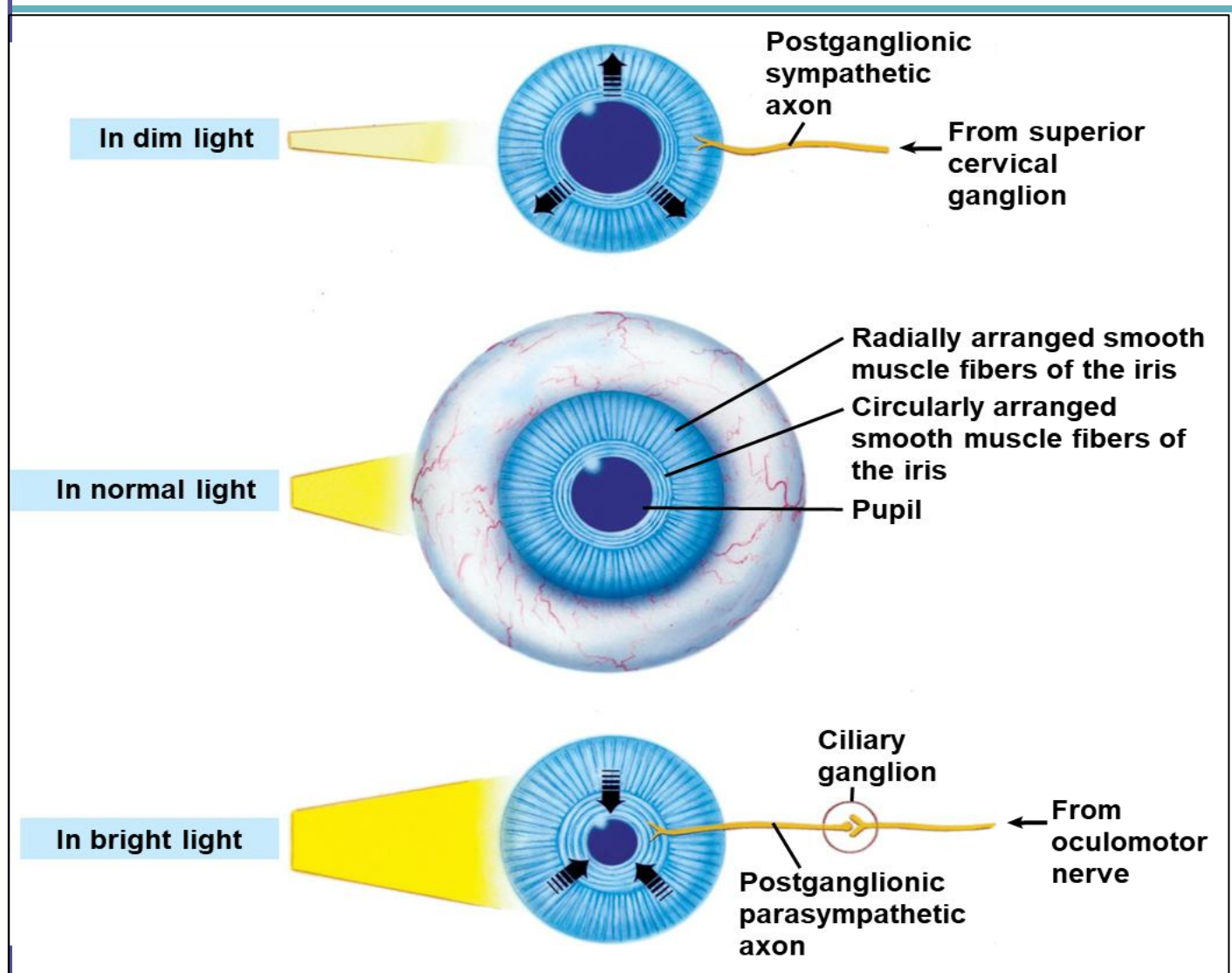


# Pupil and Iris: Dilation and Constriction of the Pupil

A. The iris can increase or decrease the diameter of the pupil.

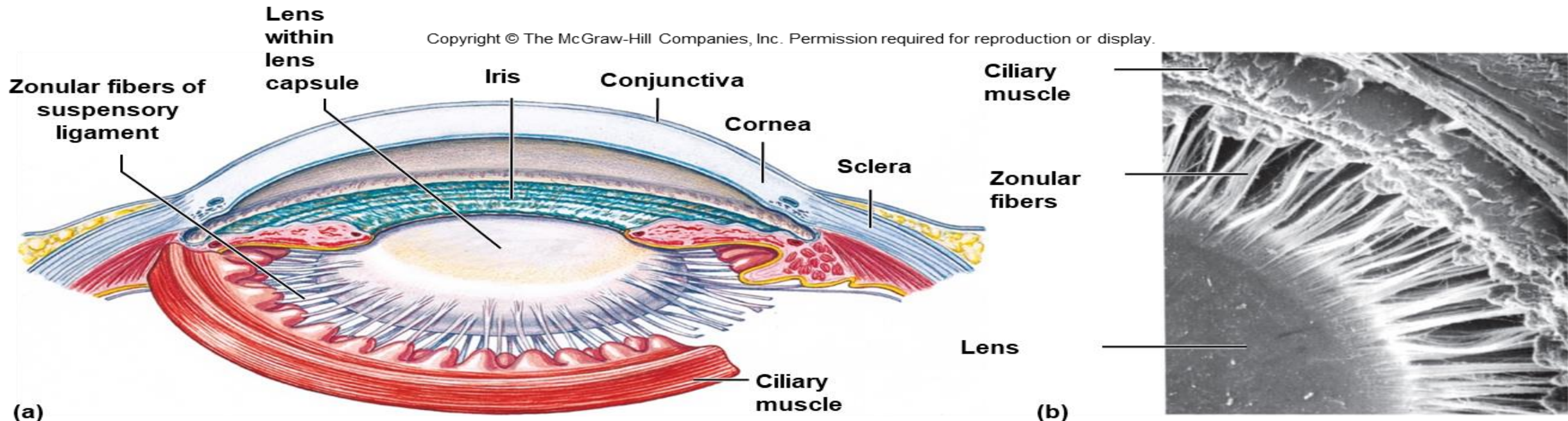
- 1) **Constriction:** contraction of circular muscles via parasympathetic stimulation
- 2) **Dilation:** contraction of radial muscles via sympathetic stimulation

B. The iris also has pigmented epithelium for eye color



# The relationship between the ciliary muscle and lens

- A. **Composed** of layers of living cells that are normally completely clear
- B. **Avascular**
- C. **Cell metabolism** is very low and anaerobic
- D. Attached to muscles called ciliary bodies and suspended from suspensory ligaments



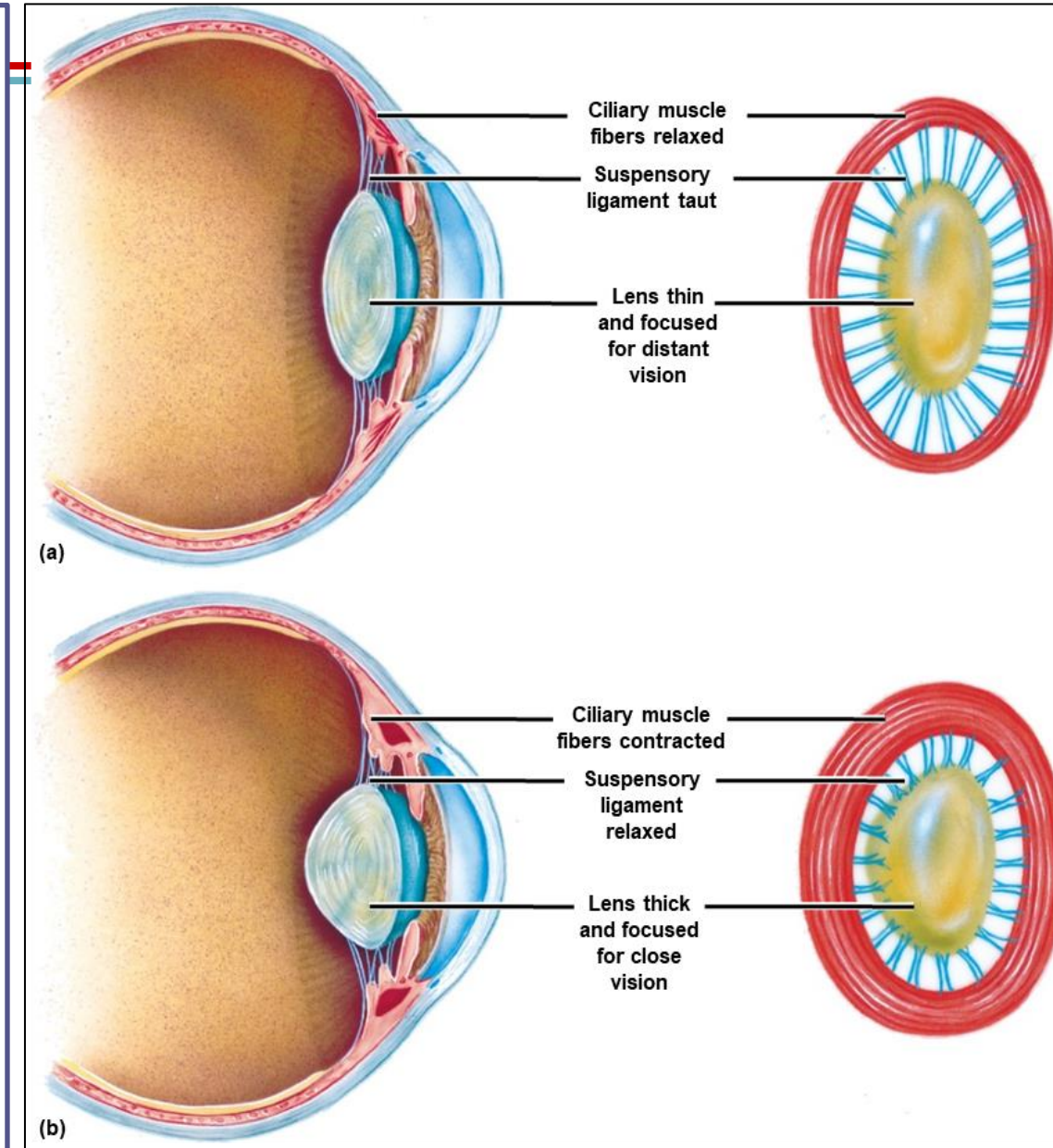
b: Courtesy of Patricia N. Farnsworth, University of Medicine and Dentistry

# Lens Accommodation

- **Accommodation** is the ability of the lens to keep an object focused on the retina as the distance between the eye and the object moves.

**A. Contraction** of the ciliary muscle allows the suspensory ligaments to relax and the lens to thicken and roundup; this is good for close vision.

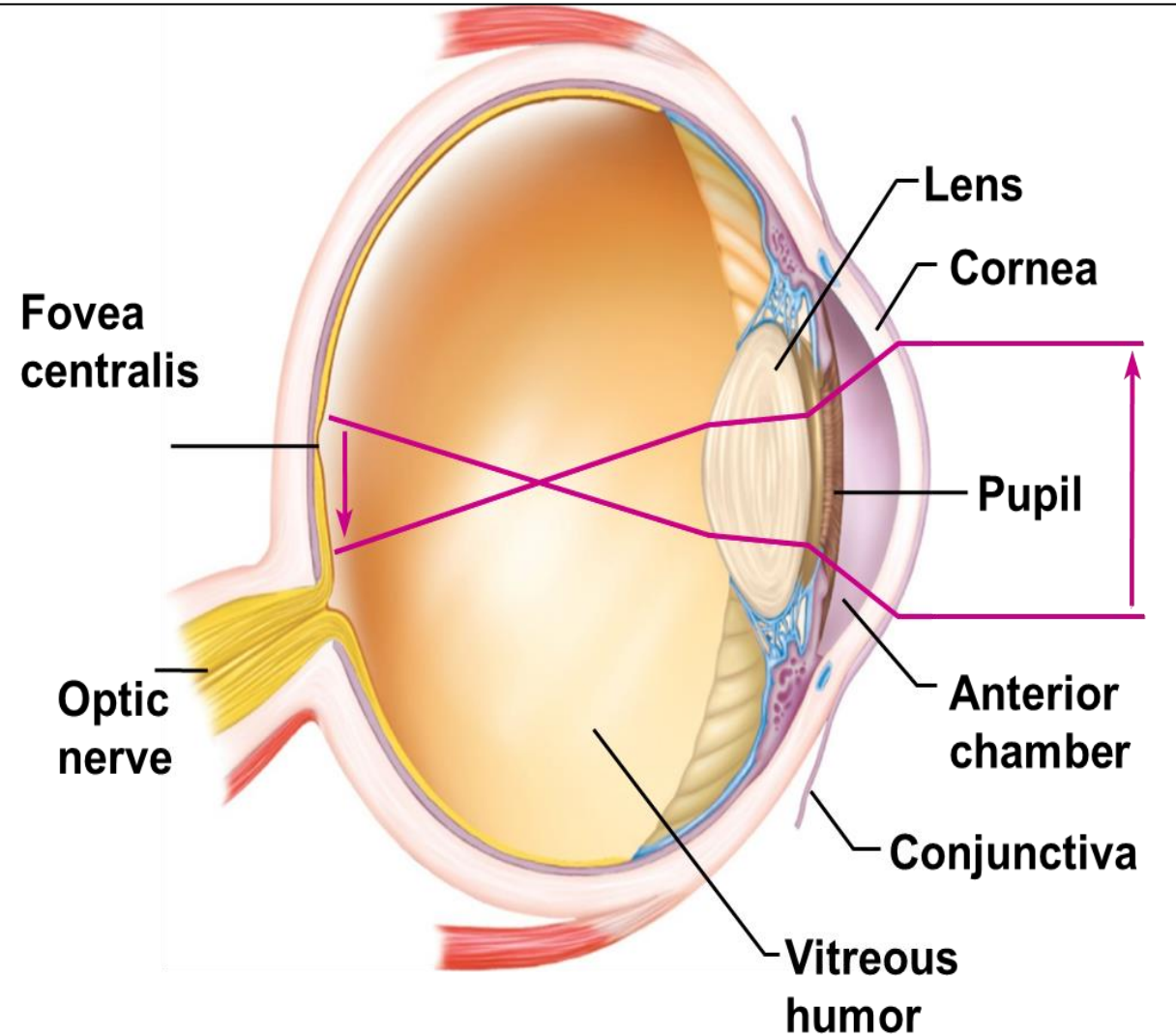
**B. Relaxation** of the ciliary muscle pulls on the suspensory ligaments, causing the lens to thin and flatten; this is good for distant vision.





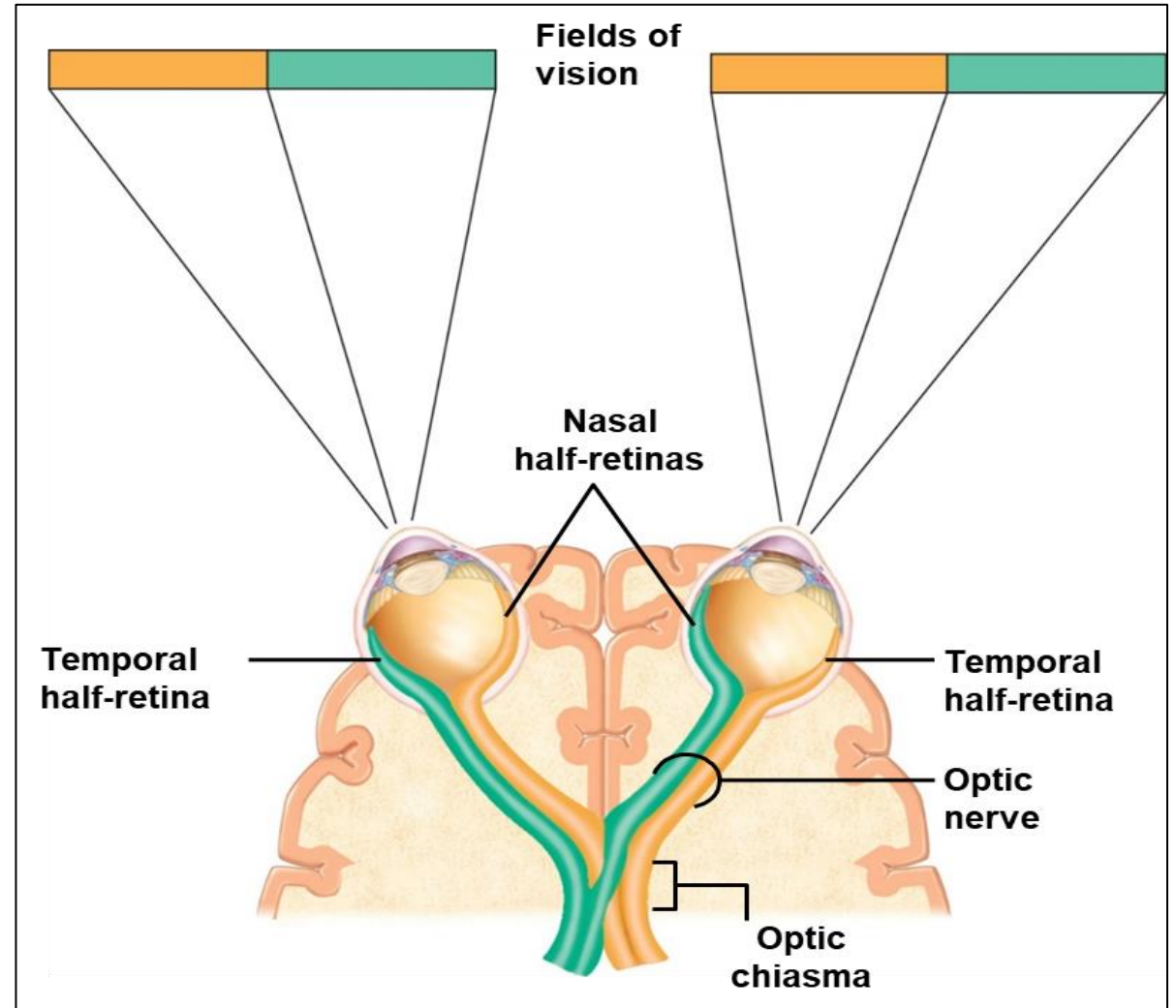
# Light Refraction

- When light passes from one medium to another with a different density, it bends.
  - a. Curvature at the point of refraction can also bend light.
  - b. Greatest refraction is at the air-cornea interface
  - c. Changing the curvature of the lens allows fine control of focus.
  - d. The humors will also contribute to the refraction
  - e. The image is flipped upside down and right to left in this process.



# Visual Fields

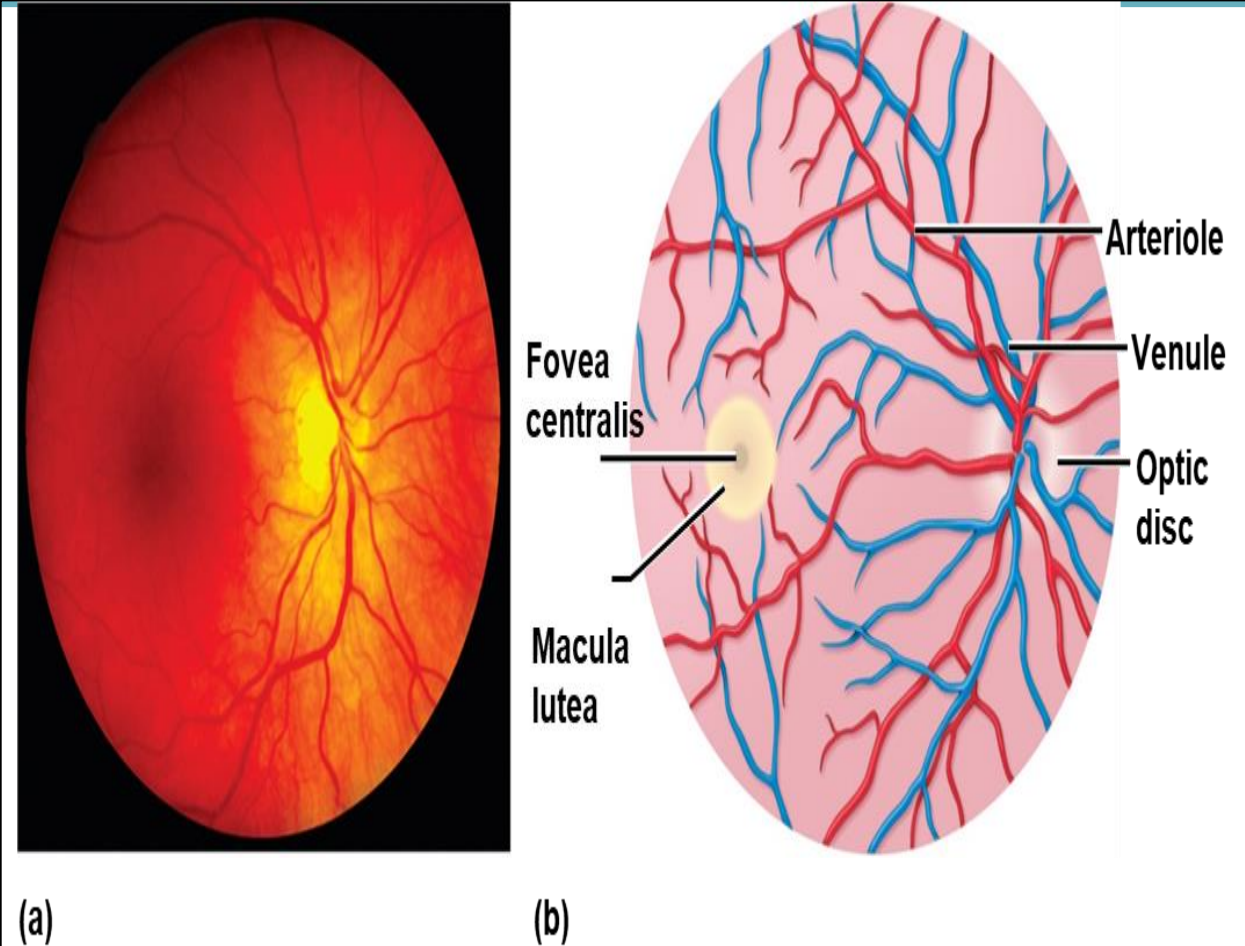
- A. **Visual fields are** the part of the external world projected onto the retina.
- B. **The right side is** projected onto the left side of the retina.
- C. **The left side is** projected onto the right side of the retina.





# The Retina

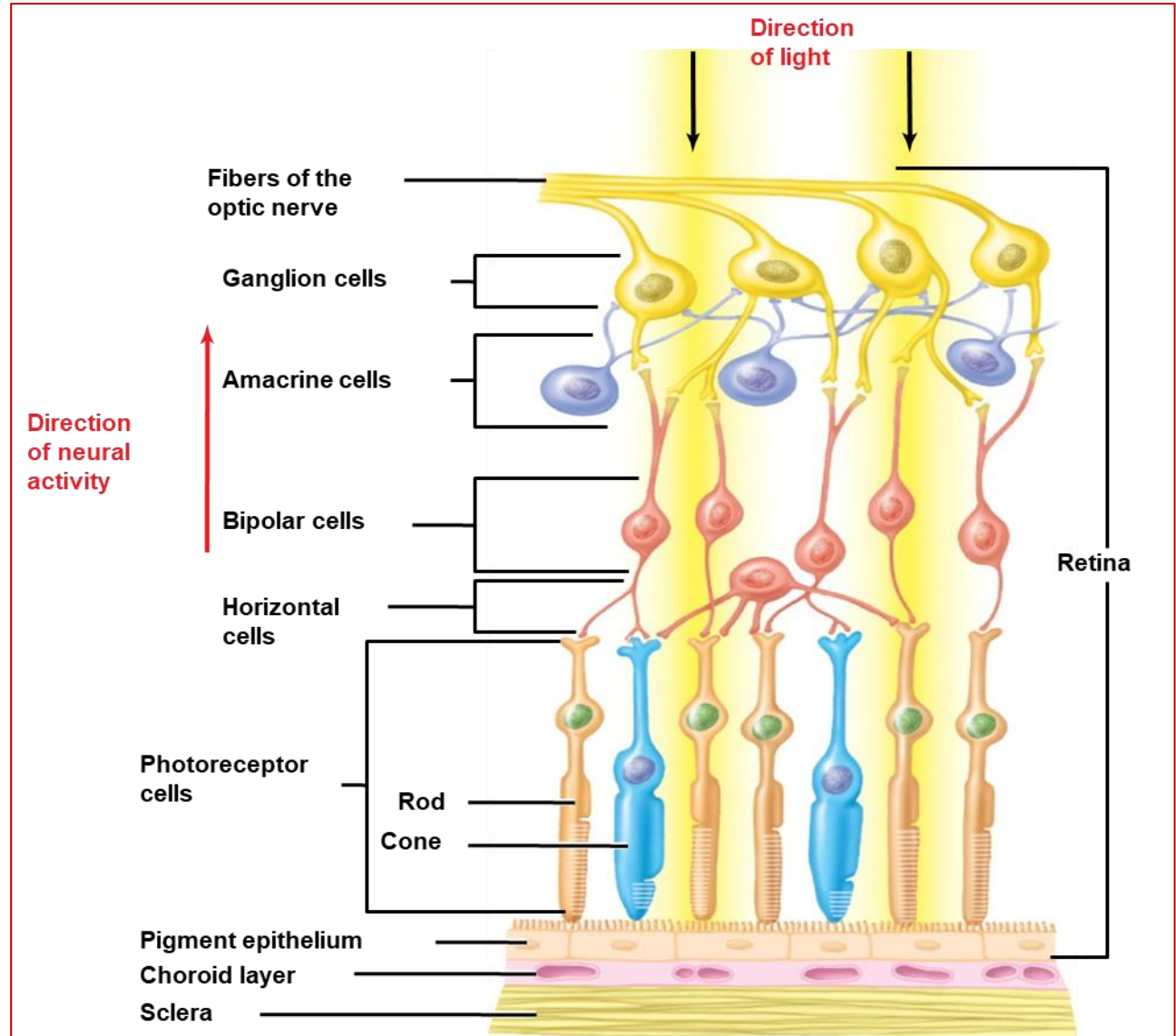
1. **The retina is a forward** extension of the brain, so the neural layers face outward toward the incoming light.
2. **Neuron axons** in the retina are gathered at a point called the optic disc (blind spot) and exit as the optic nerve.
3. **Blood vessels** also enter and leave here.



A view of the retina with an ophthalmoscope

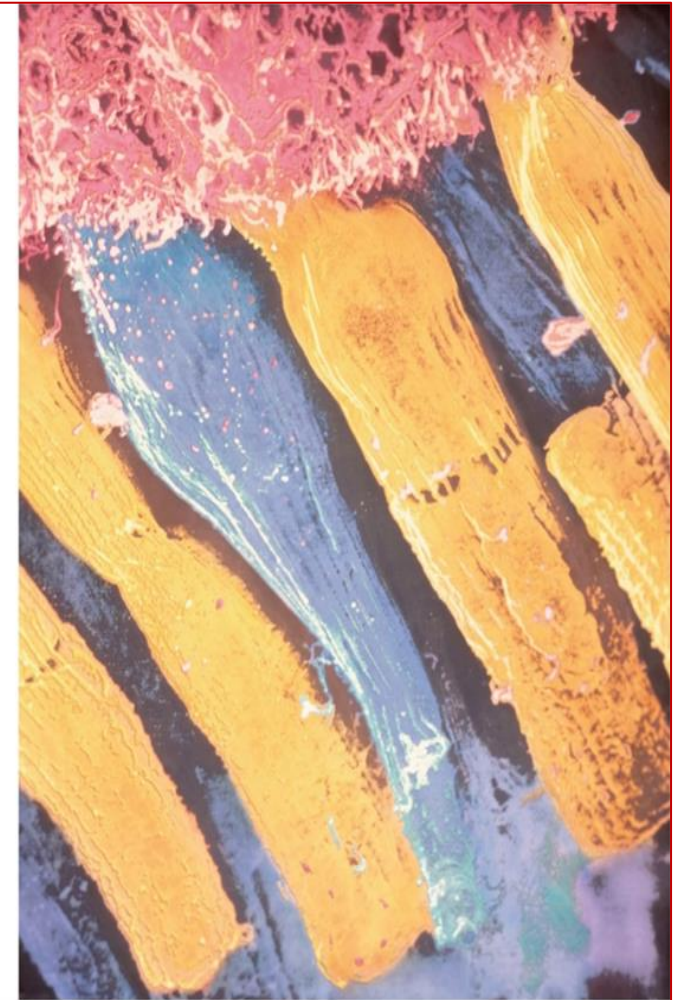
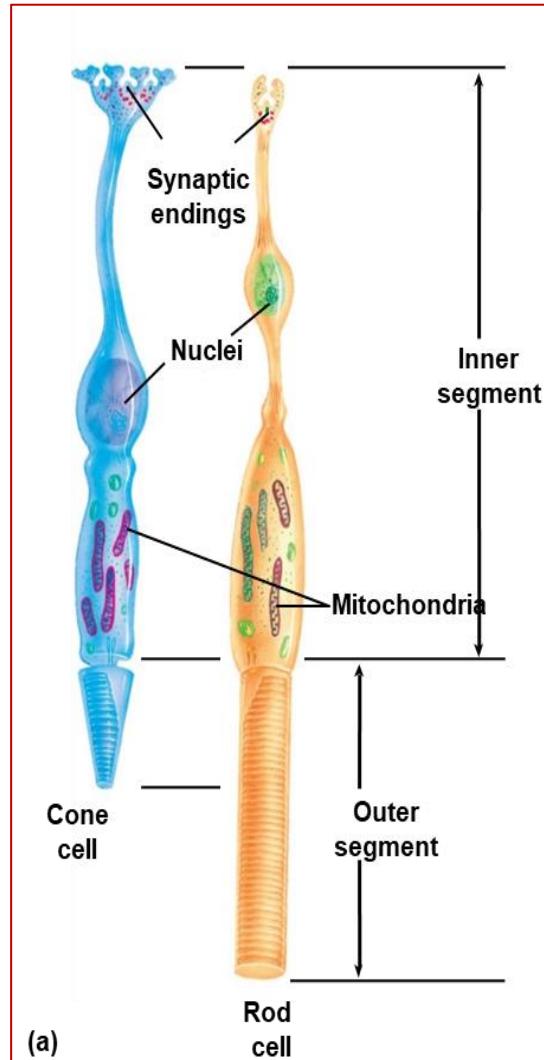
# Layers of the Retina

1. **Photoreceptors** (rods and cones) are in the inner layer (toward the vitreous body).
2. **These synapse** on a middle layer of bipolar cells, which synapse on the outer layer of ganglion cells.
3. **There are also** horizontal cells and amacrine cells within the layers



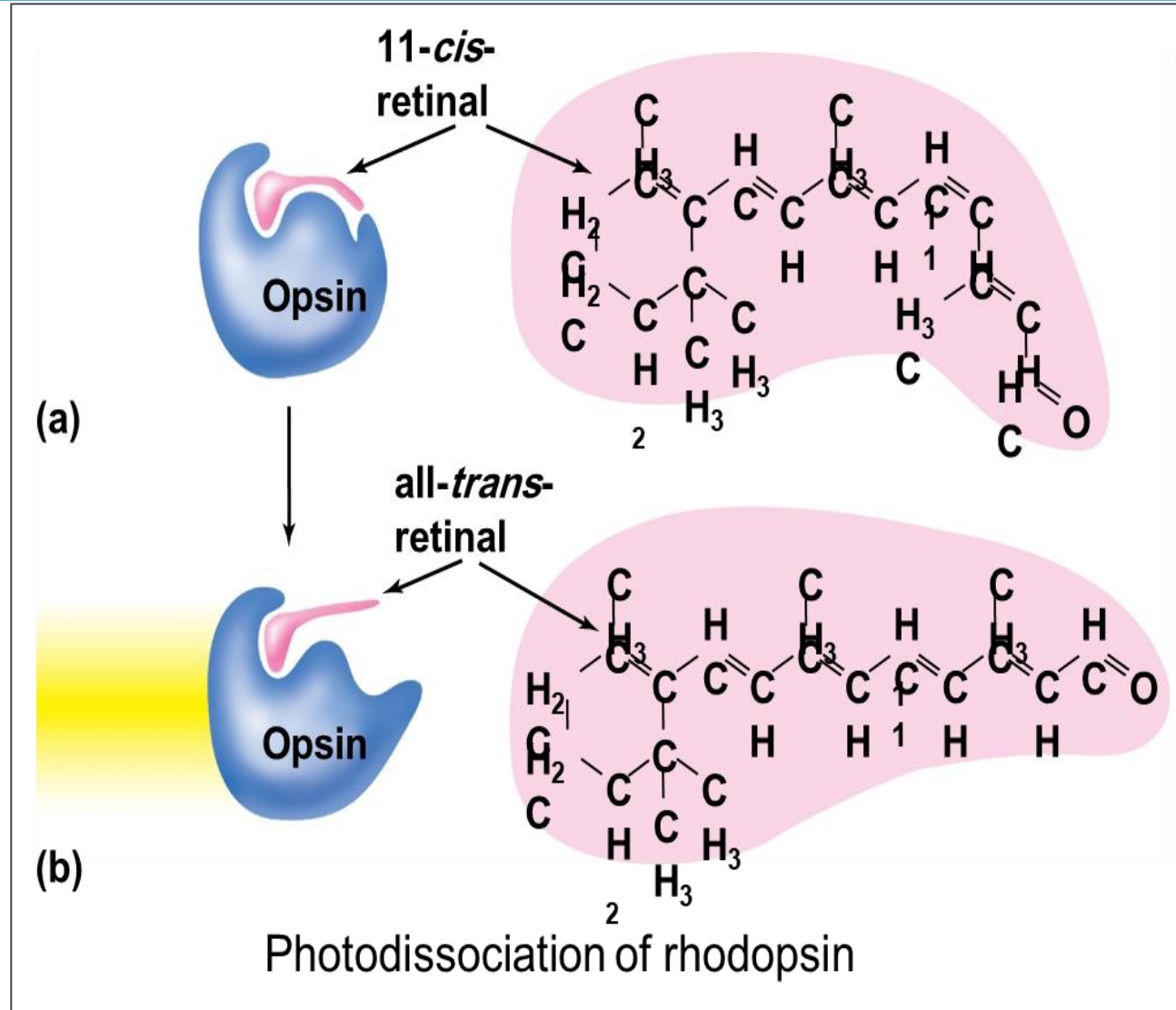
# Rods and Cones

- Consist of:
  - Outer segment**; full of flattened discs with photopigment molecules.
  - Inner segment** that contains the cell organelles



# Effects of light on the rods

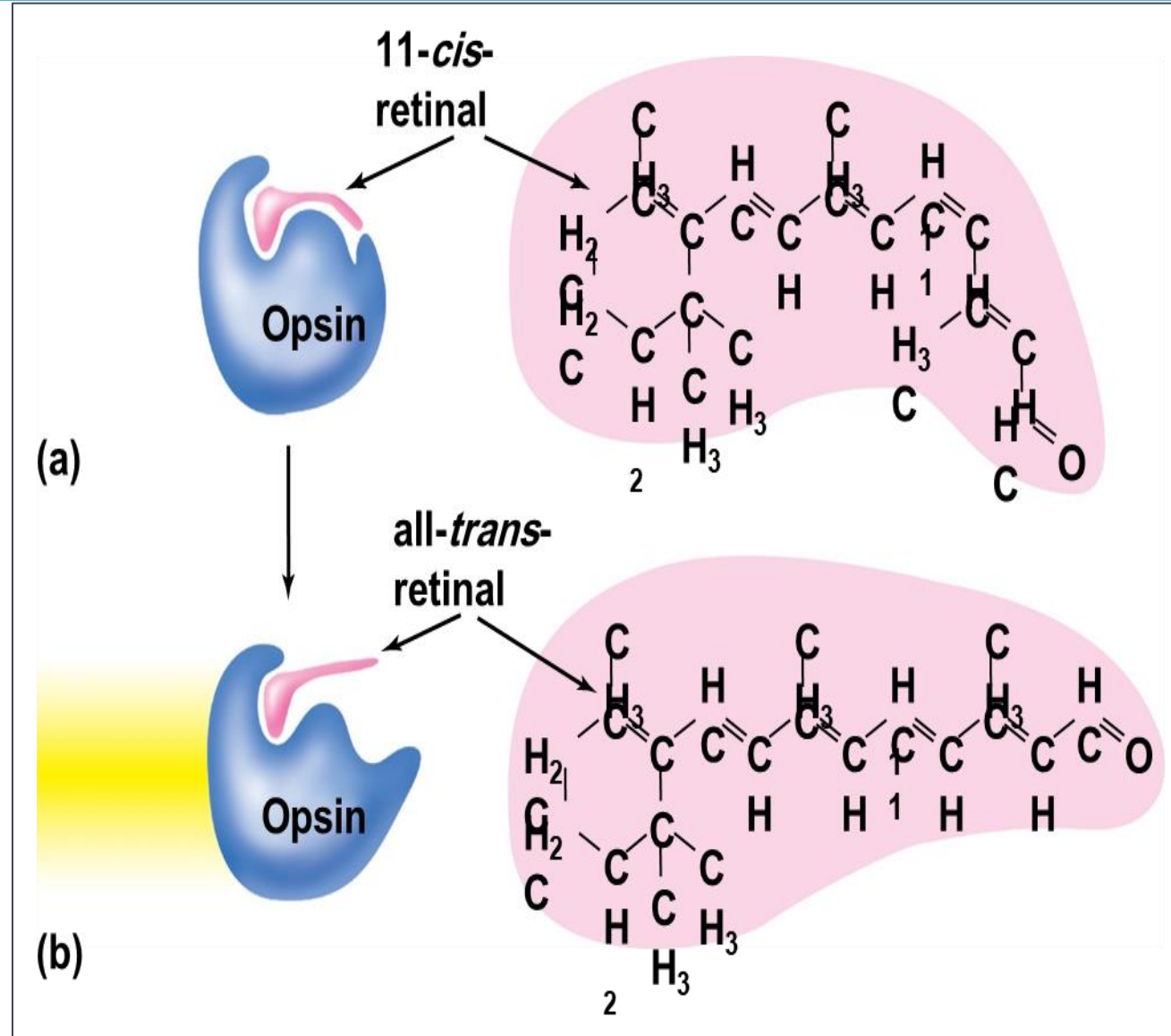
1. **Rods allow** black-and-white vision in low light.
2. **Contain the purple** pigment rhodopsin, which absorbs green light best.
  - a. **Absorption** causes rhodopsin to dissociate into retinaldehyde and opsin.
  - b. **Retinaldehyde** (also called retinal) is derived from vitamin A.
  - c. **Called the bleaching reaction**





# Visual Cycle of Retinal

- a. **In rhodopsin**, retinal exists in an *11-cis* form.
- b. **After bleaching**, the retinal is in an *all-trans* form.
- c. **To be reincorporated** into retinal, it must be converted back into *11-cis*.
- d. **This occurs** in the pigment epithelial cells.





# Dark Adaptation

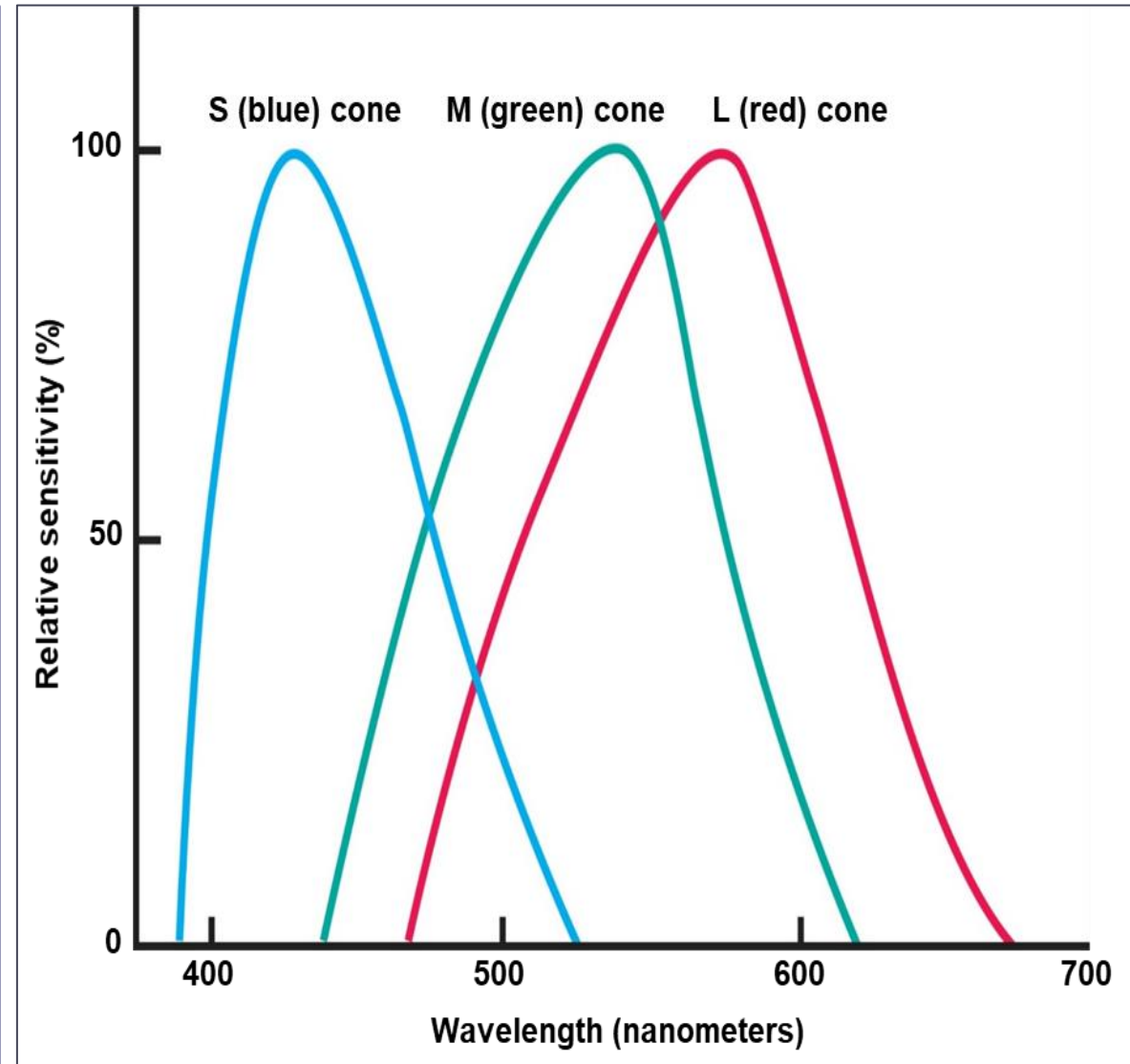
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A. When a person enters a dark room after being in the light, there are fewer photopigments in the rods and cones.

B. After about 20 minutes, more visual pigments are produced, and the person's eyes adapt to the dark.

# Cones and color vision

1. **Cones are** less sensitive to light but allow color vision and greater visual acuity.
2. **Trichromatic vision** involves three types of cones.
  - a. S: short wavelengths, blue
  - b. M: medium wavelengths, green
  - c. L: long wavelengths, red
3. **Instead of opsins**, photopigments have photopsins with retinene.
4. **Photopsins** vary in each type of cone.
5. **Cone response** depends on wavelength and the intensity of the light



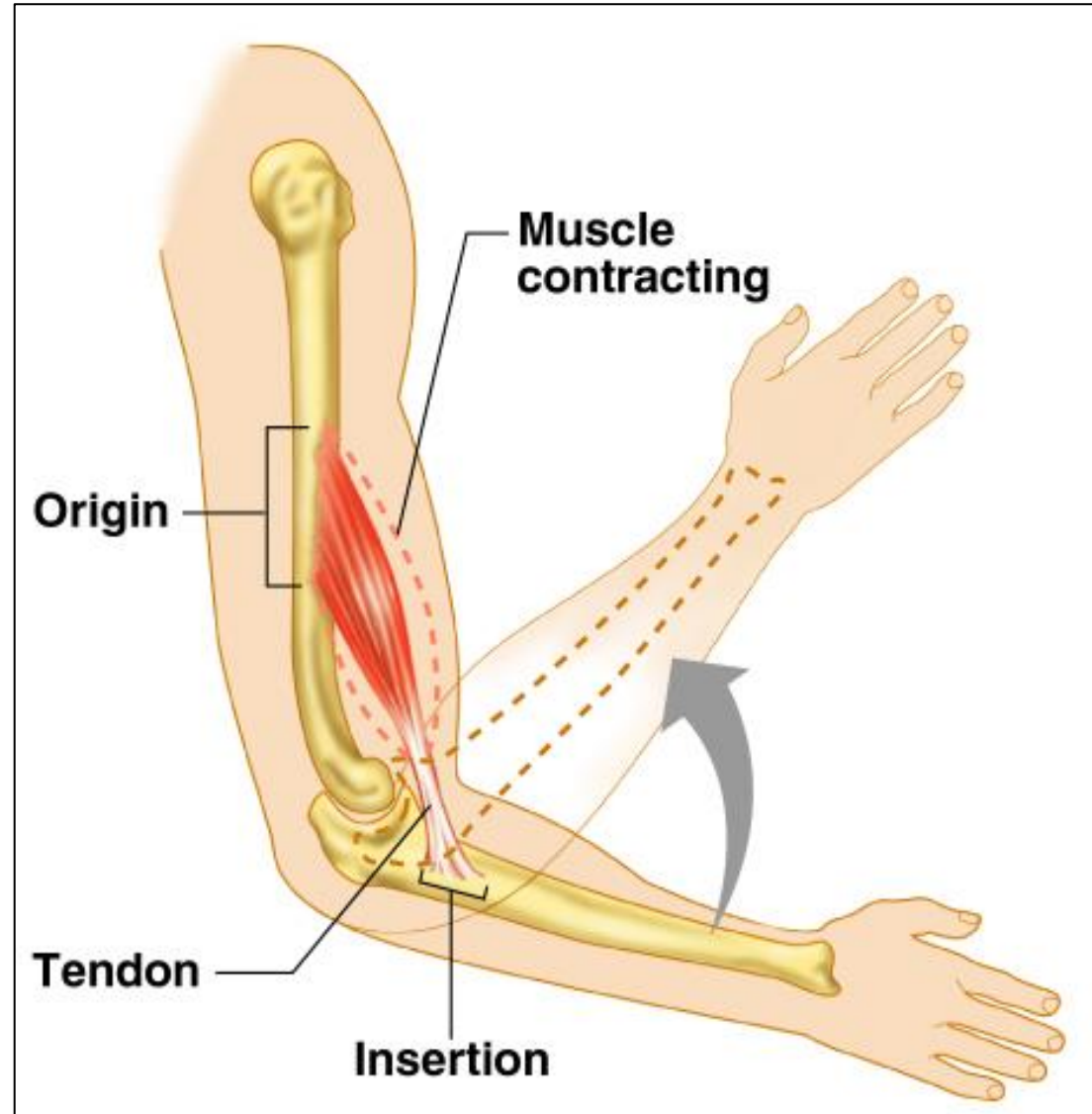
# **Chapter 12 Muscle: Mechanisms of Contraction and Neural Control**

# Muscles

- Muscles are responsible for all types of body movement.
- Three basic muscle types are found in the body
  - A. Skeletal muscle:** attaches to bones, Voluntary movement (body movement, facial expression).
  - B. Cardiac muscle:** heart wall, Involuntary heart movement (Pumping blood).
  - C. Smooth muscle:** Walls of hollow internal organs (wall of internal viscera Stomach, Intestines, Bladder, Uterus, Blood vessels), Involuntary movement.

# Skeletal Muscles

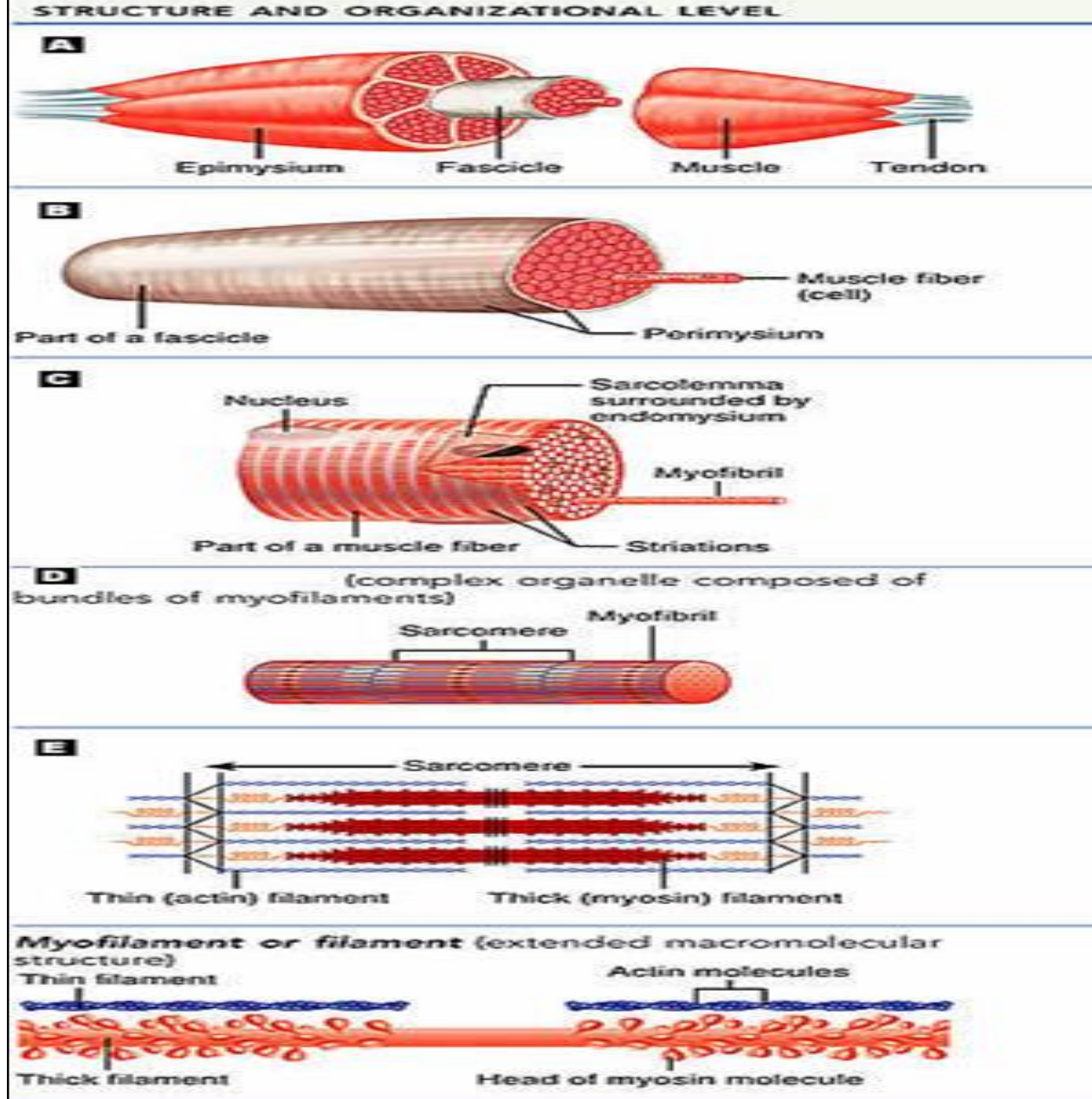
1. **Generally**, both ends of a muscle are attached to bone by tough tendons.
2. **When a muscle** contracts, it shortens.
  - A. **This places** tension on tendons connecting it to a bone.
  - B. **This moves** the bone at a joint.





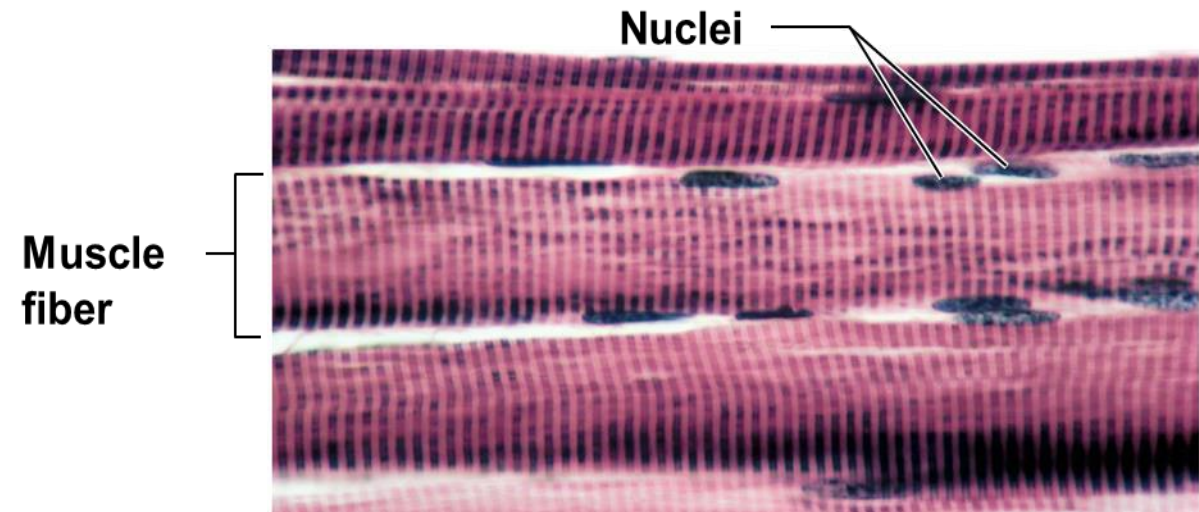
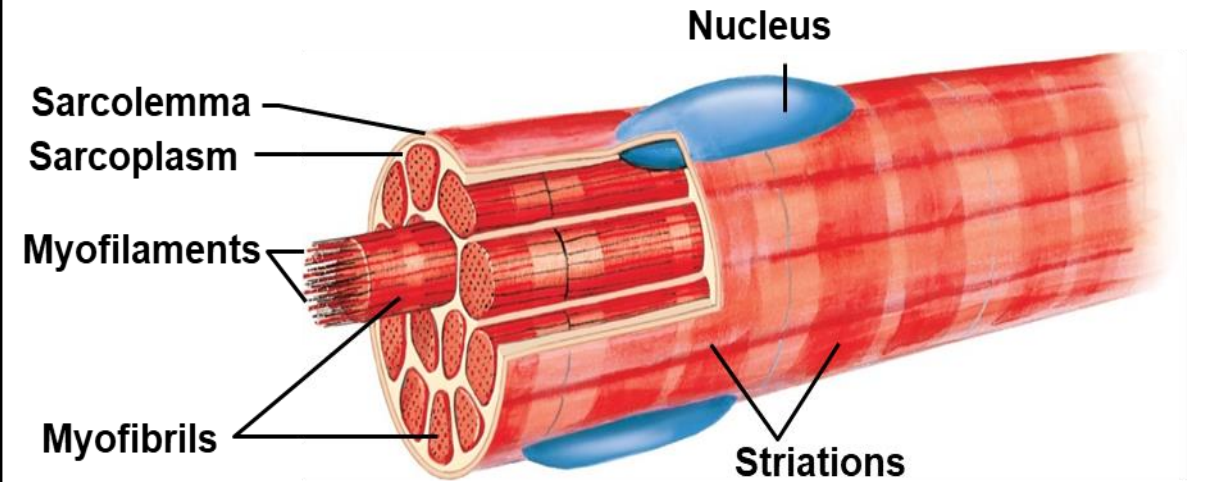
# Structure of Skeletal Muscles

- **Connective** tissue subdivides the muscle into fascicles.
- **Each** fascicle is subdivided into muscle fibers (myofibers).



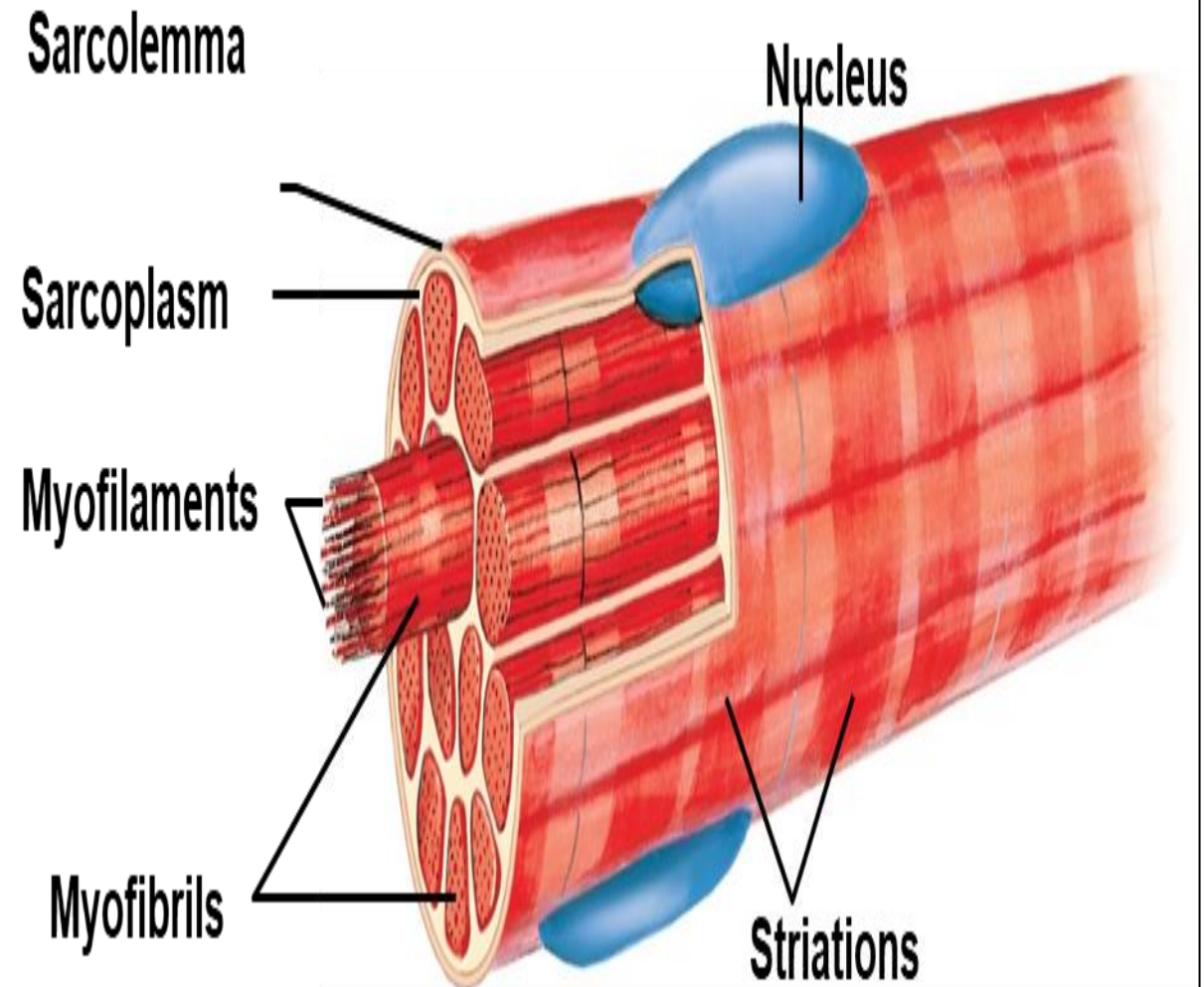
# Muscle Fiber Structure (Muscle cell)

- A. Have many of the organelles found in other cells.
- B. Have plasma membranes called sarcolemma.
- C. Are multinucleated; form a syncytium.
- D. Are striated
  - 1) I bands: light bands
  - 2) A bands: dark bands



# Muscle Fiber Banding

- Each fiber has densely packed subunits called **myofibrils** that run the length of the muscle fiber.
  - A. **Stacked in register** so that the dark and light bands align.
  - B. **Composed** of **thick** and **thin myofilaments**





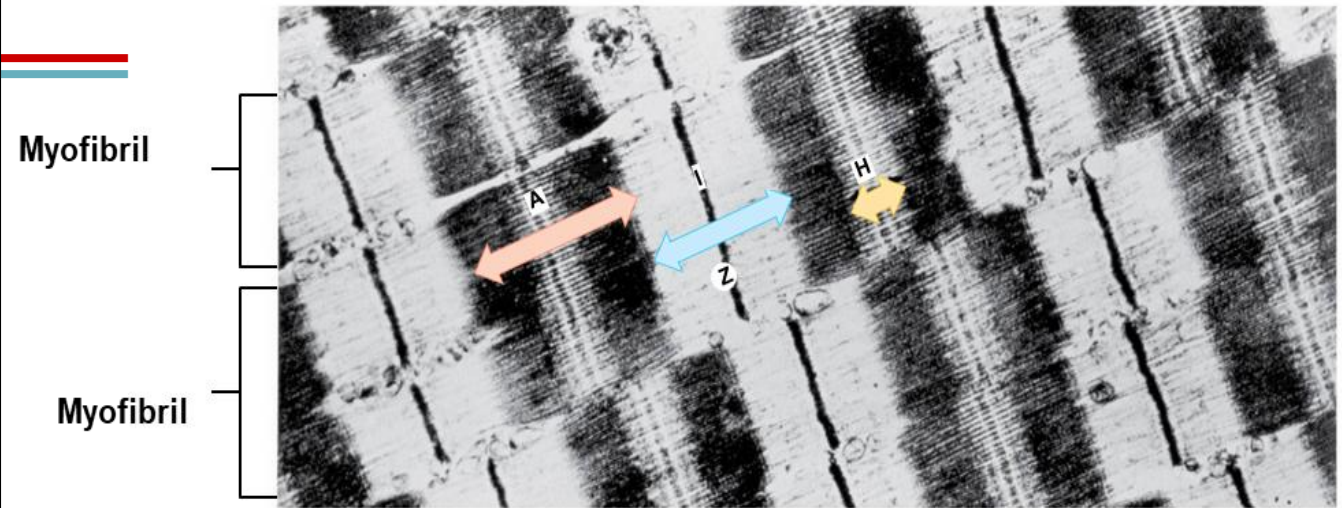
# Striations

- Produced by **thick** and **thin filaments**

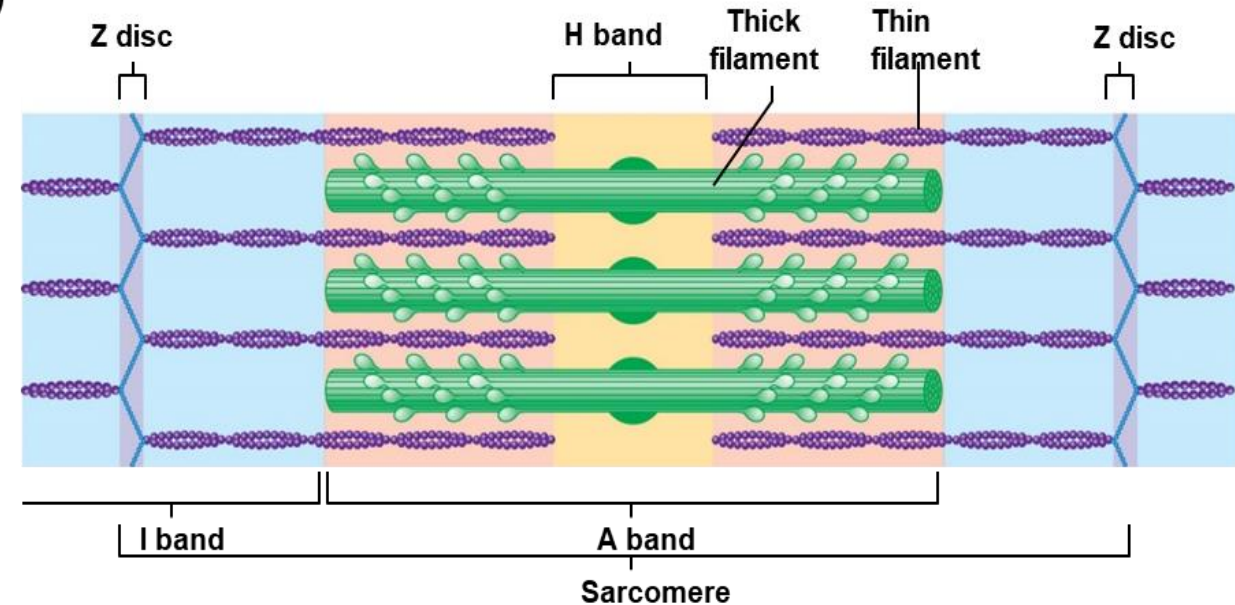
- I bands** contain only thin filaments, primarily of the protein, actin
- A bands** contain all of the thick filament with some thin filament overlap; the thick filament is the protein, myosin
- H bands** are the center of the A band with no thin filament overlap.
- Z discs** (lines) are found in the center of each I band.

- The sarcomere:

- The basic** subunit of striated muscle contraction.
- Area from** one Z disc to the next
- Titin:** protein that runs from the Z disc to the M line and allows elastic recoil.
- M lines** are found in the center of each A band and help hold down thick filaments.
- In three dimension**, the sarcomere forms a hexagonal pattern.

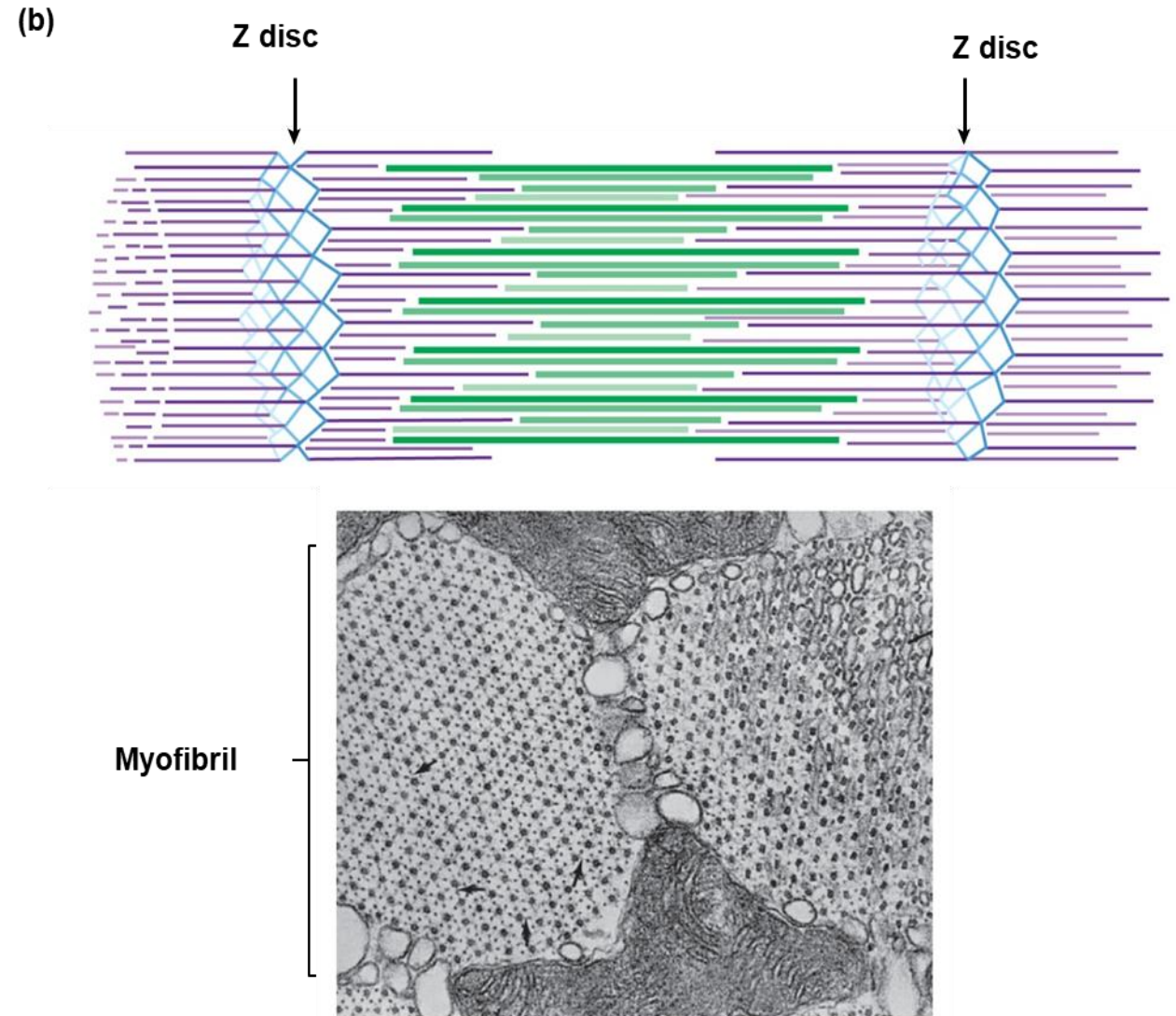
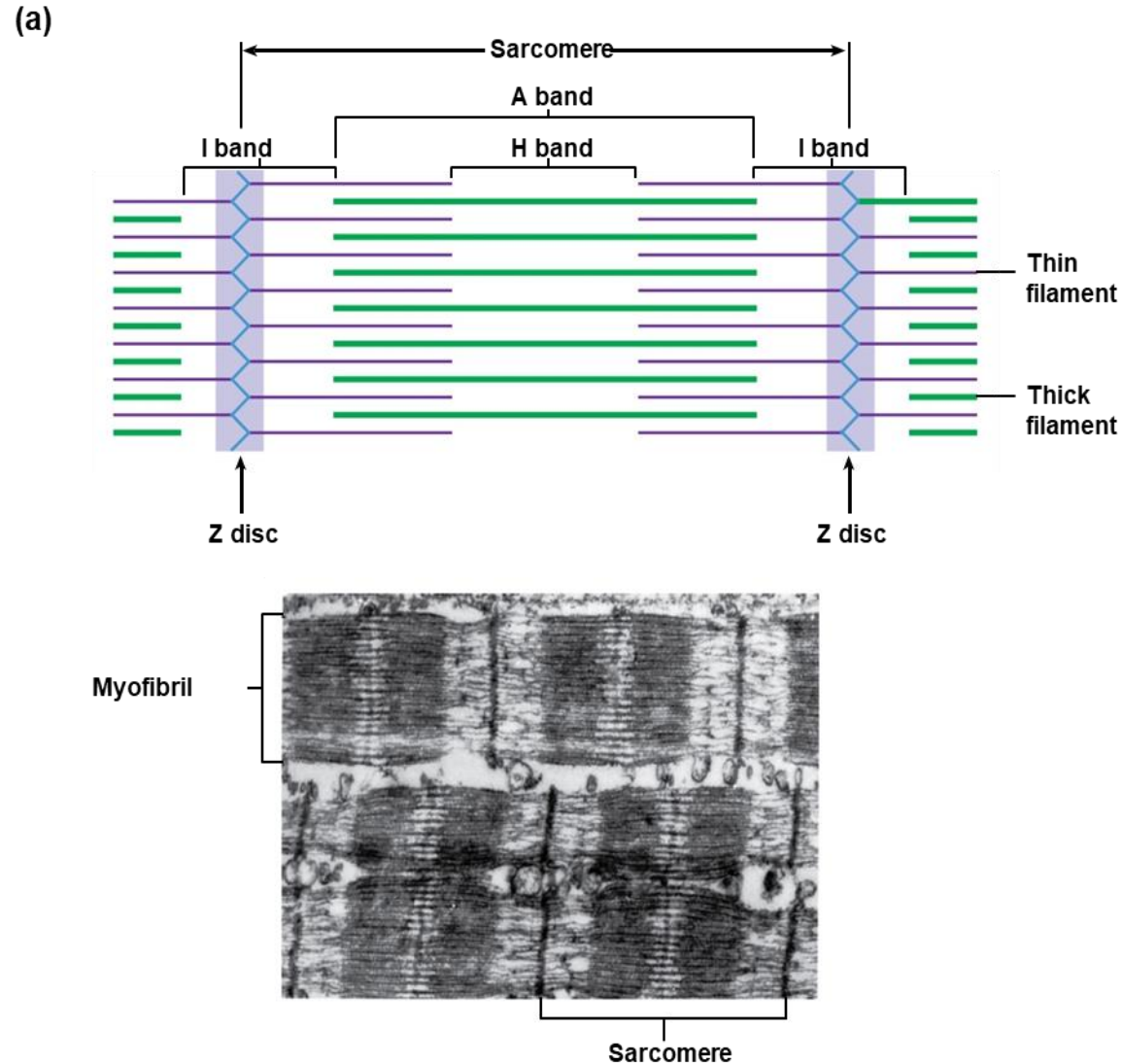


(a)



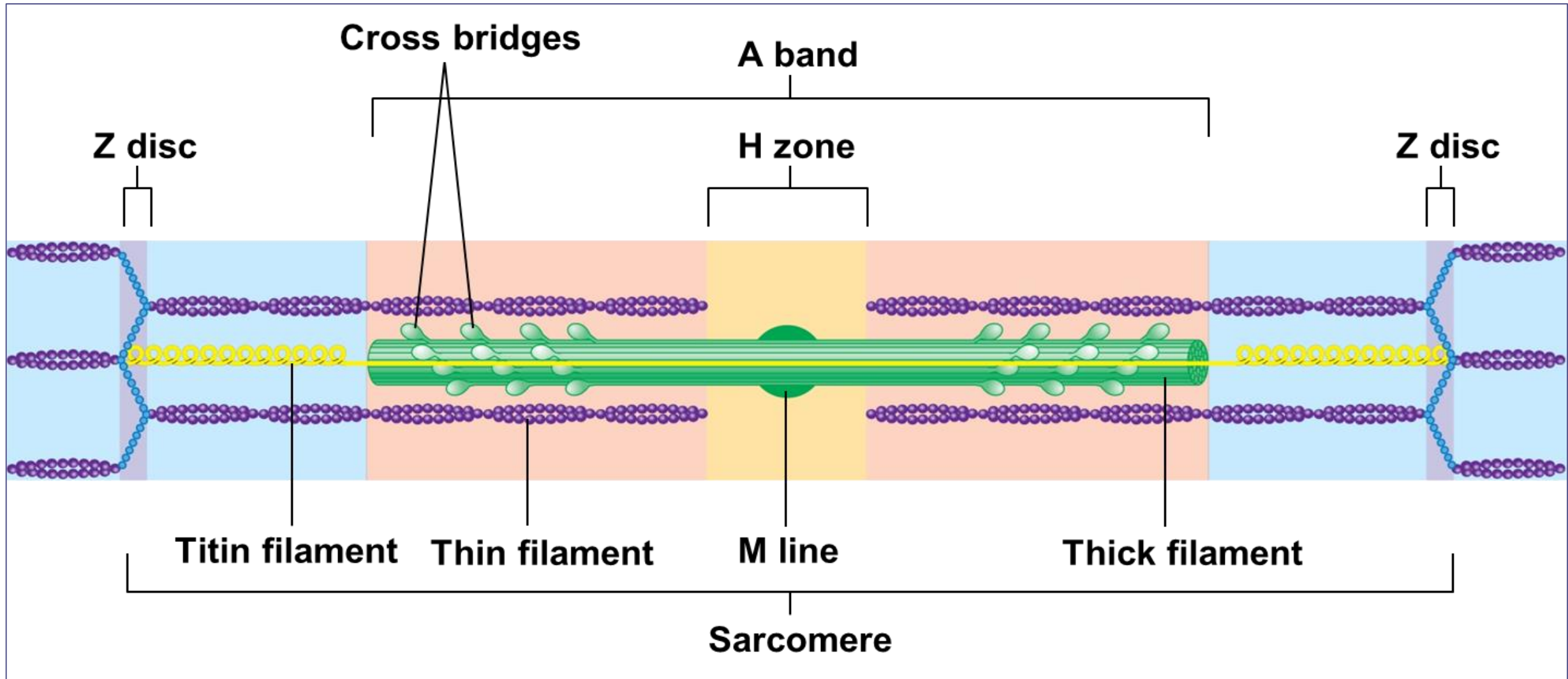
(b)

# Arrangement of Thick and Thin Filaments





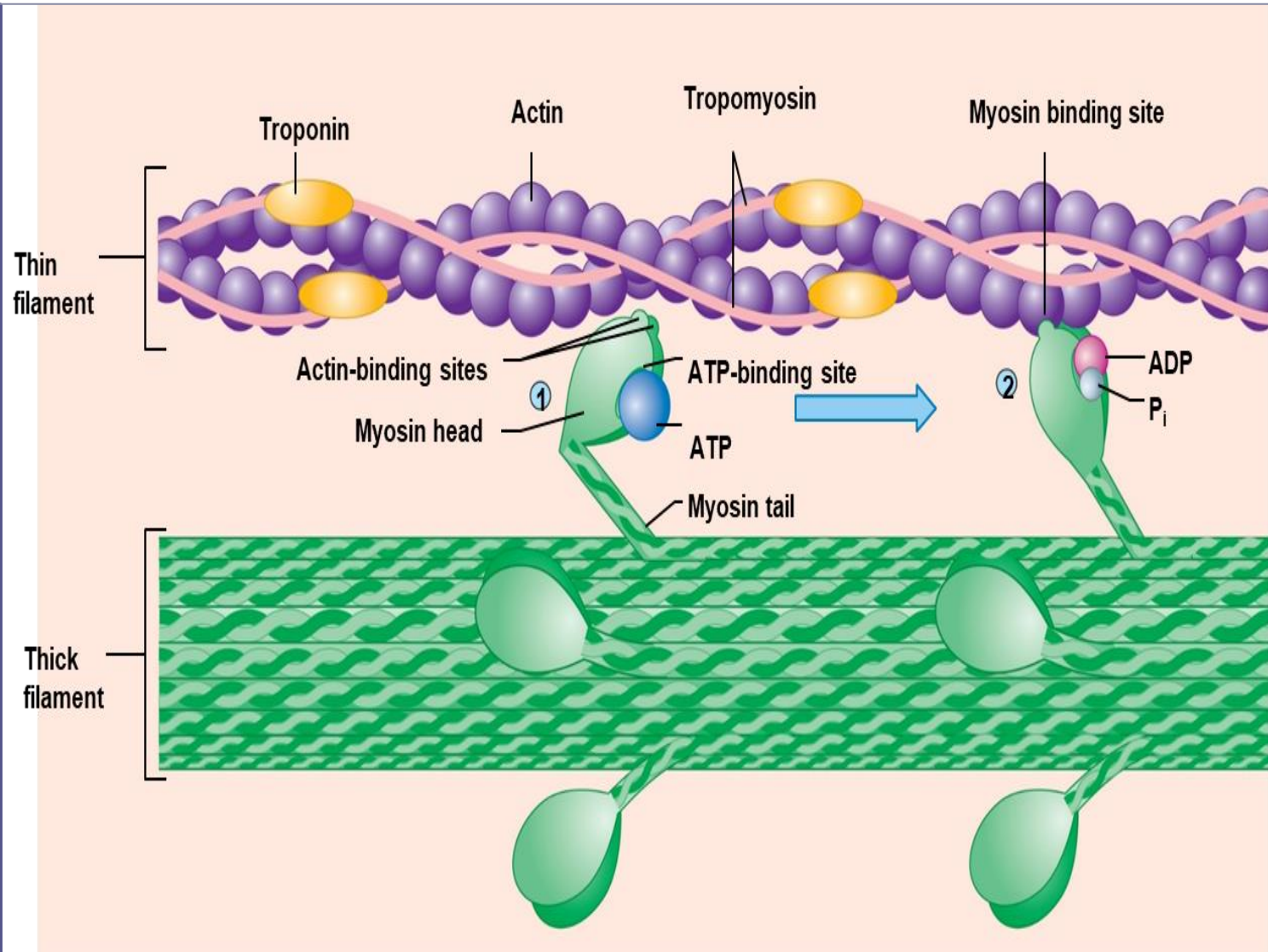
# Titin Filaments and M Lines



# Cross bridges

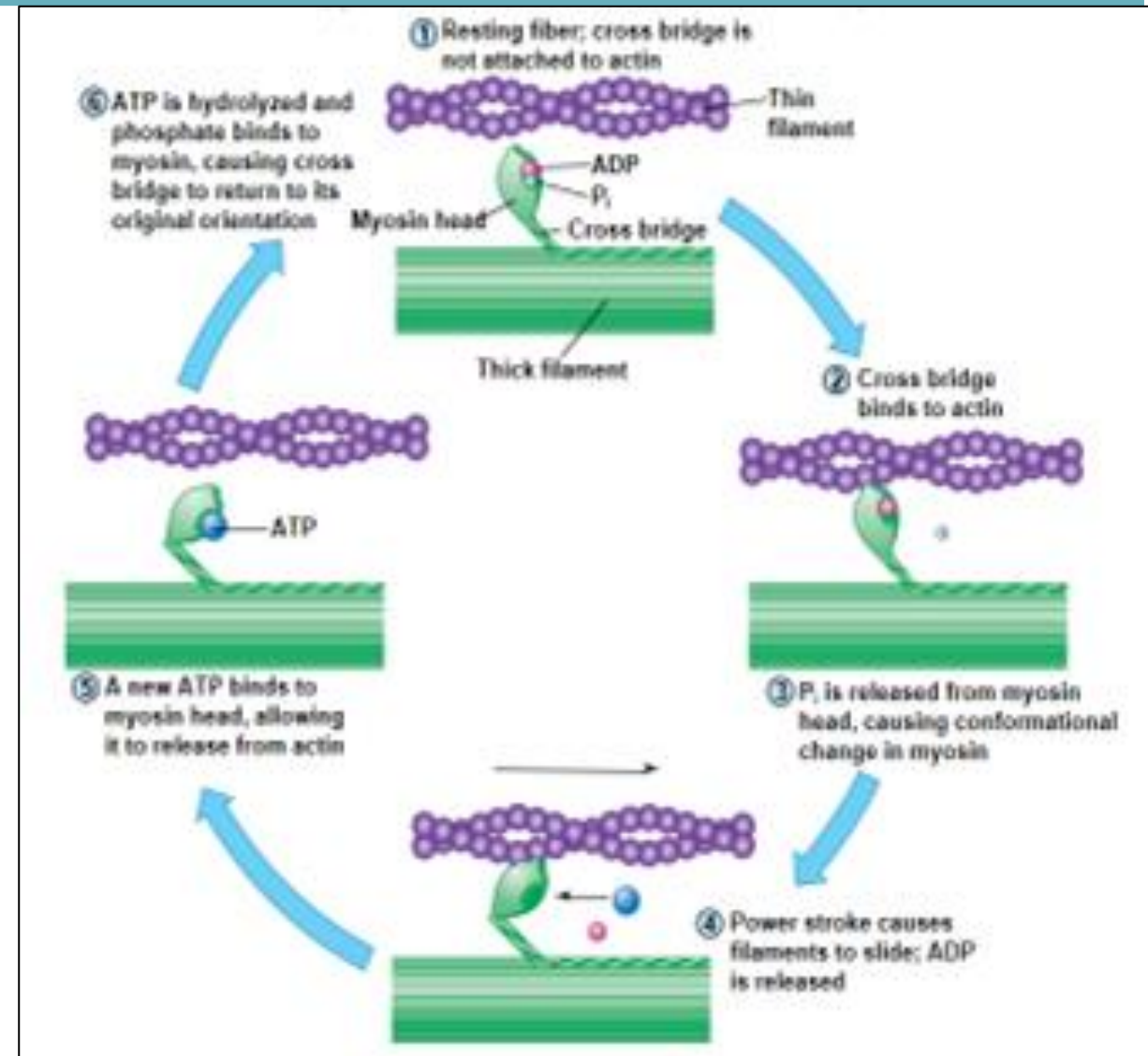
More about myofilaments:

1. **Thick:** composed of the protein myosin.
  - **Each protein** has two globular heads with actin-binding sites and ATP-binding sites.
1. **Thin:** composed of the protein actin.
  - **Have proteins** called tropomyosin and troponin that prevent myosin binding at rest.



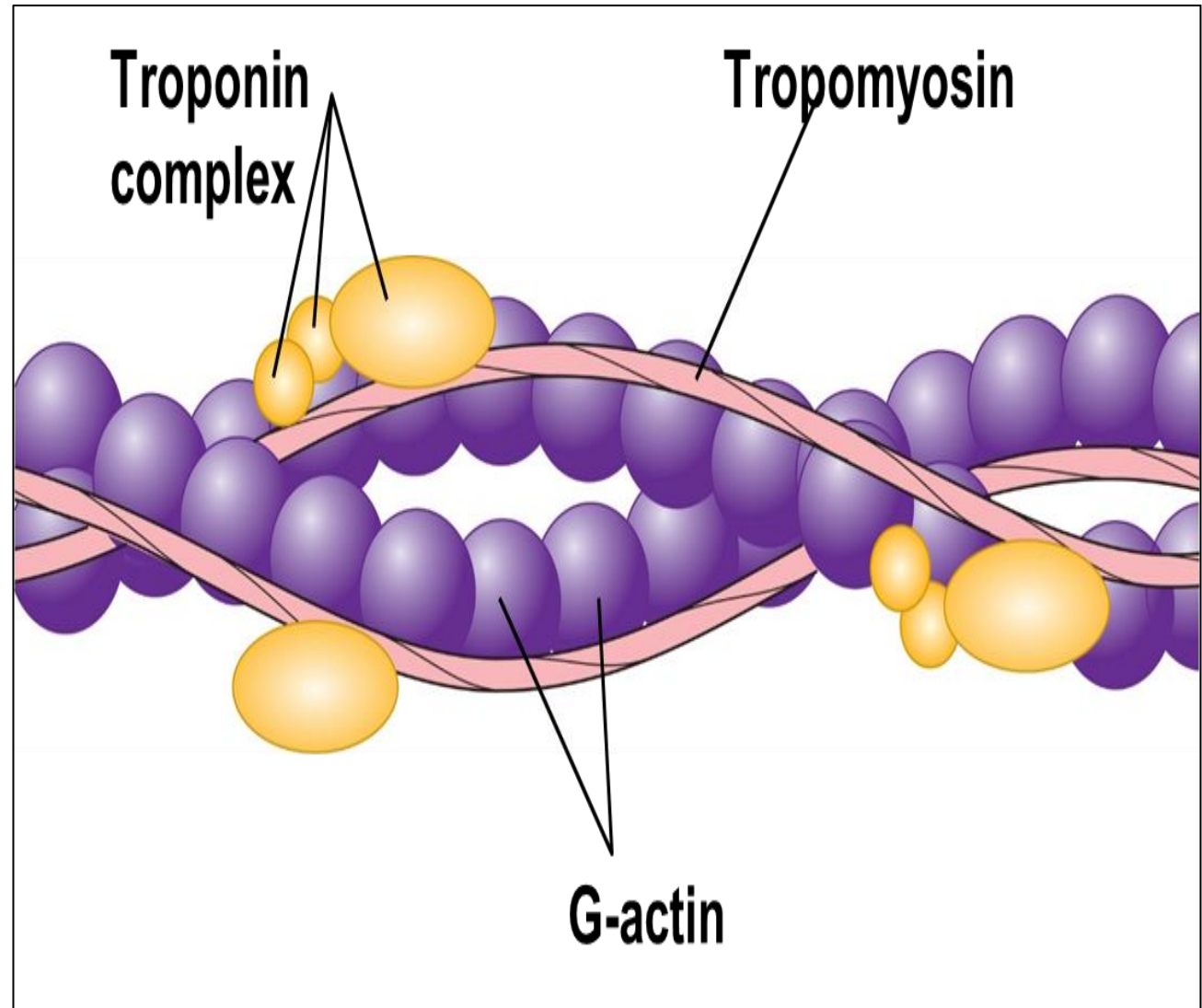
# Action of sliding

1. **Sliding** is produced by several cross bridges that form between myosin and actin.
  - A. The myosin head serves as a myosin ATPase enzyme, splitting ATP into ADP +  $P_i$ .
  - B. This allows the head to bind to actin when the muscle is stimulated.
2. **Release of  $P_i$  upon binding** cocks the myosin head, producing a power stroke that pulls the thin filament toward the center.
3. **After the power stroke**, ADP is released and a new ATP binds.
  - A. This makes myosin release actin.
  - B. ATP is split.
4. **The myosin** head straightens out and rebinds to actin farther back.
5. **Continues until** the sarcomere has shortened



# Regulation of Contraction

1. **F-actin** is made of 300-400 G-actin subunits, arranged in a double row and twisted to form a helix.
2. **Tropomyosin** physically blocks cross bridges.
3. **Troponin complex**:
  - a. **Troponin I** inhibits binding of myosin.
  - b. **Troponin T** binds to tropomyosin.
  - c. **Troponin C** binds to calcium.

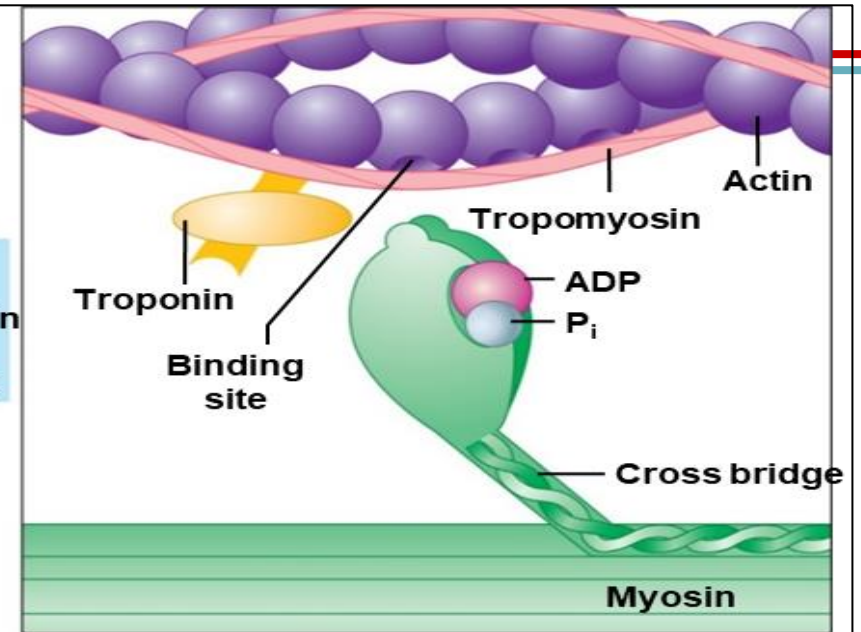




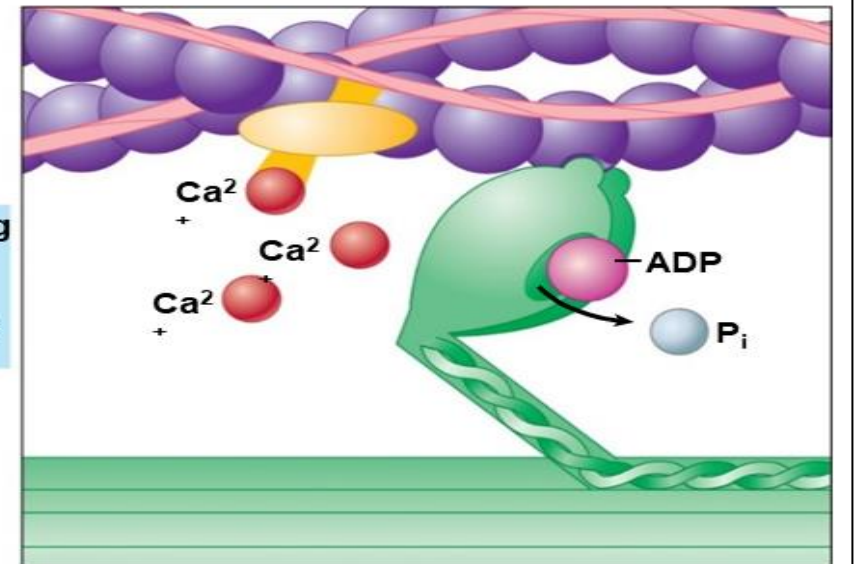
# Role of Calcium

1. **When muscle** cells are stimulated,  $\text{Ca}^{2+}$  is released inside the muscle fiber.
2. **Some attaches** to troponin C, causing a conformational change in troponin and tropomyosin.
3. **Myosin is allowed** access to form cross bridges with actin.

Relaxed muscle:  
tropomyosin  
blocks the  
binding site



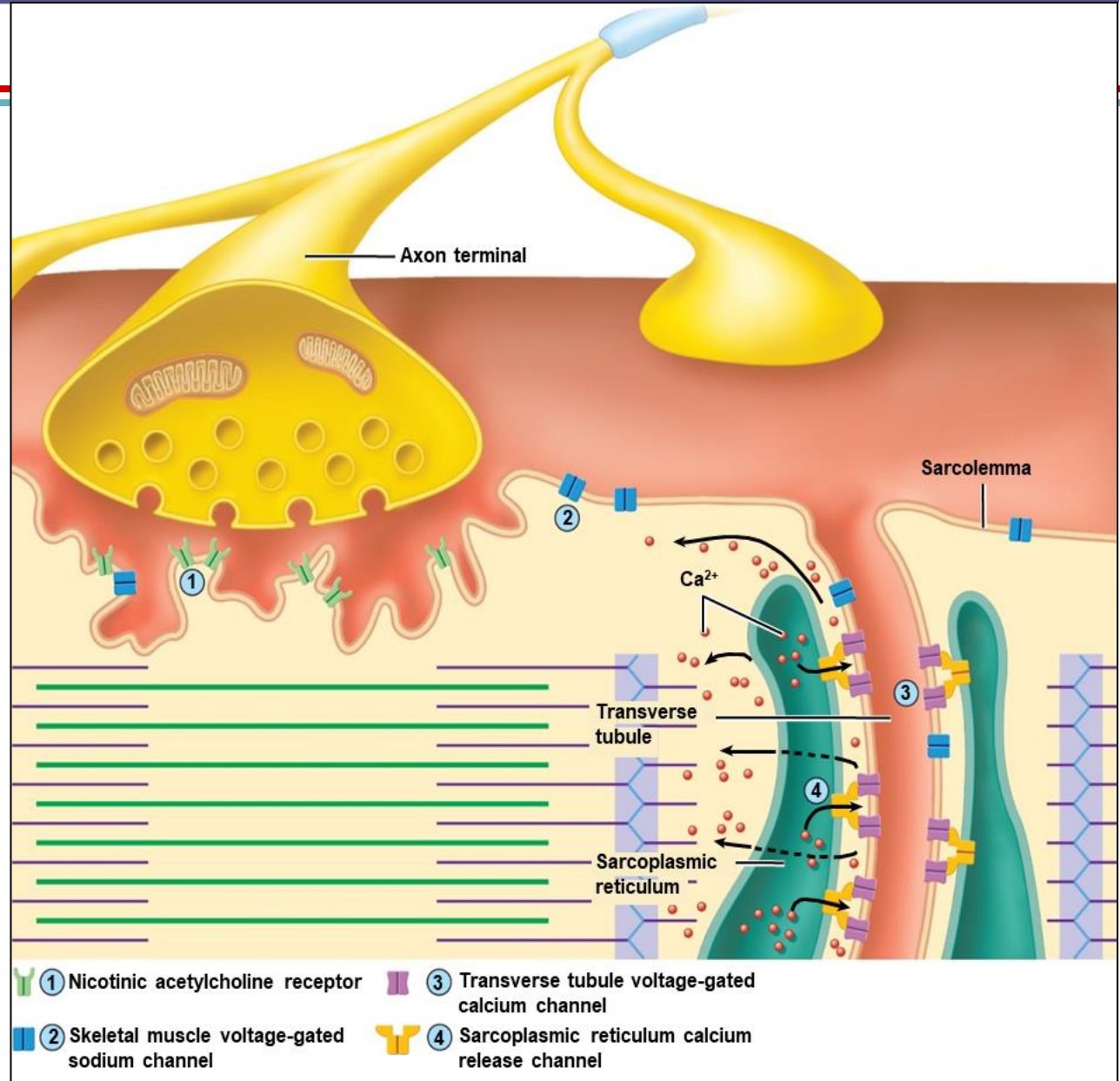
Contracting muscle:  
myosin  
head binds  
to actin





# Excitation-Contraction Coupling

- A. **Sarcoplasmic** reticulum (SR) is modified endoplasmic reticulum that stores  $\text{Ca}^{2+}$  when muscle is at rest.
- B. **When a muscle** fiber is stimulated by an action potential, Voltage-gated calcium channels in transverse tubules change shape and cause calcium channels in SR to open,  $\text{Ca}^{2+}$  diffuses out of calcium release channels (ryanodine receptors), can bind to troponin C.
- C. **At the end of a contraction**,  $\text{Ca}^{2+}$  is actively pumped back into the SR.



## Muscle Relaxation

- A. Action potentials cease.
- B. Calcium release channels close
- C.  $\text{Ca}^{2+}$ -ATPase pumps move  $\text{Ca}^{2+}$  back into SR (active transport).
- D. No more  $\text{Ca}^{2+}$  is available to bind to troponin C
- E. Tropomyosin moves to block the myosin heads from binding to actin