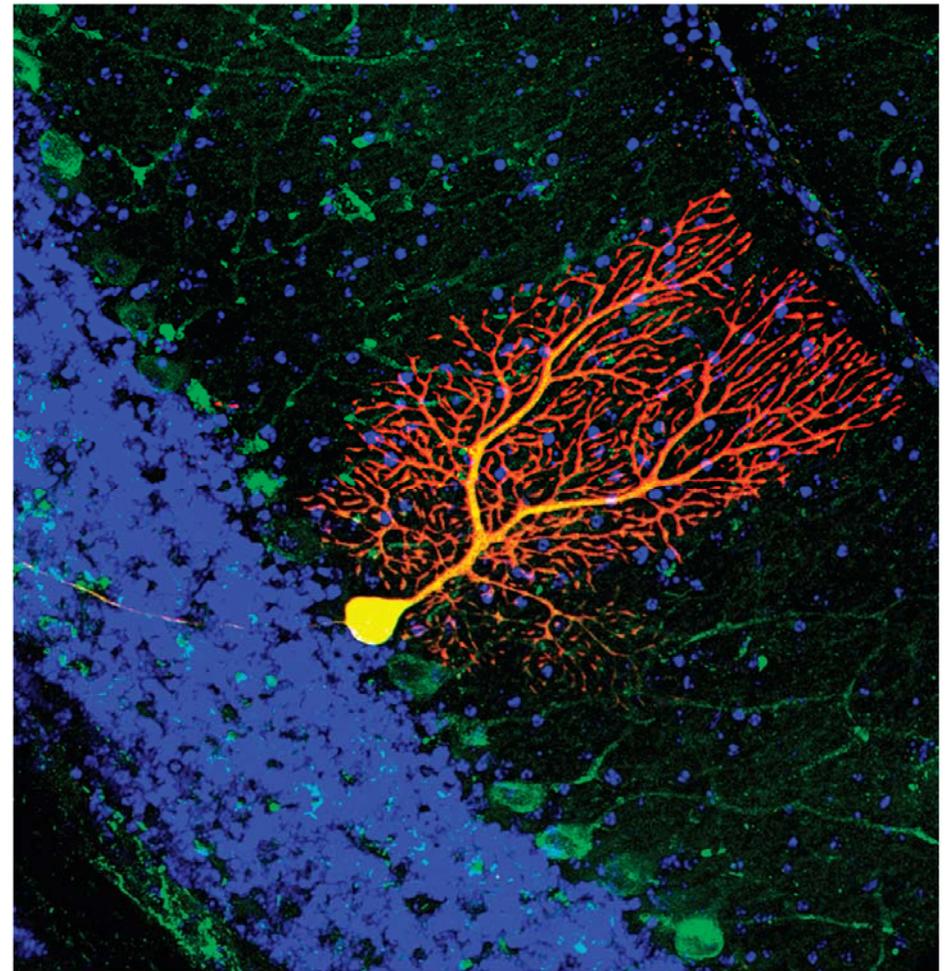


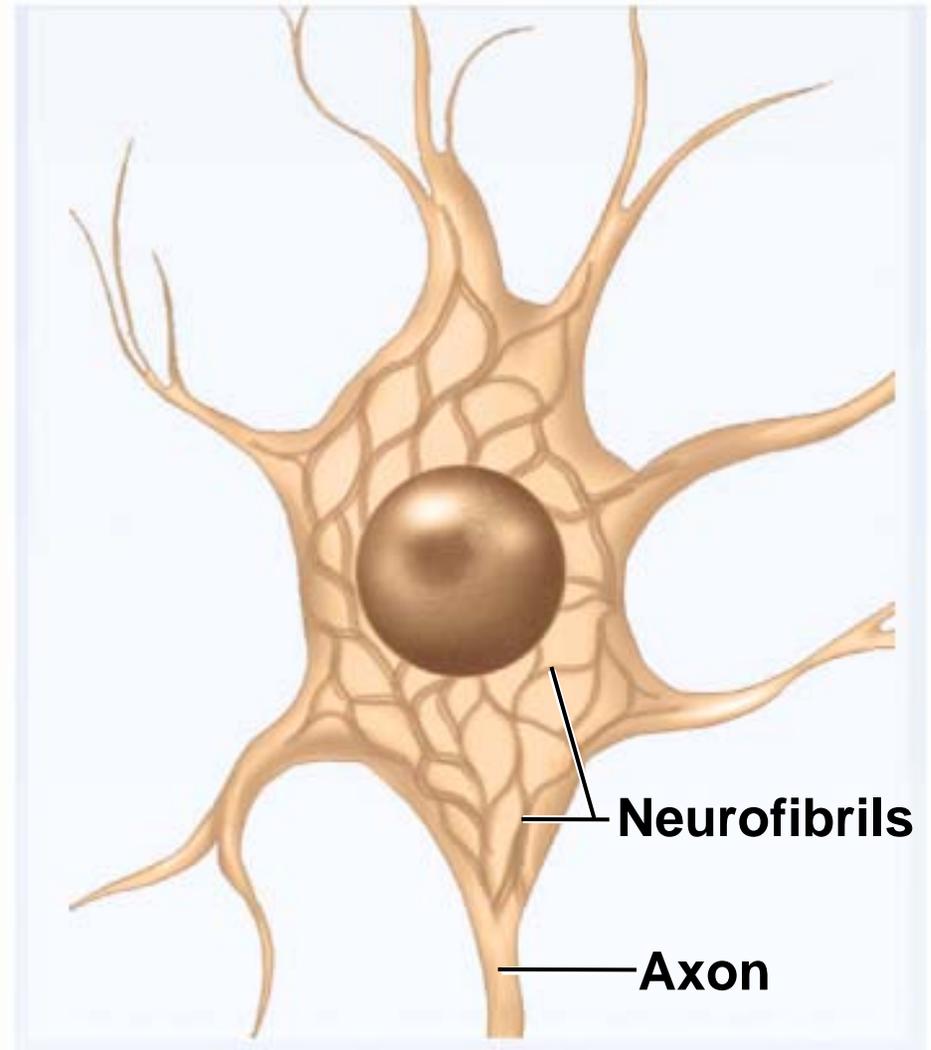
Chapter 12

Nervous Tissue



Nervous Tissue

- overview of the nervous system
- properties of neurons
- supportive cells (neuroglia)
- electrophysiology of neurons
- synapses



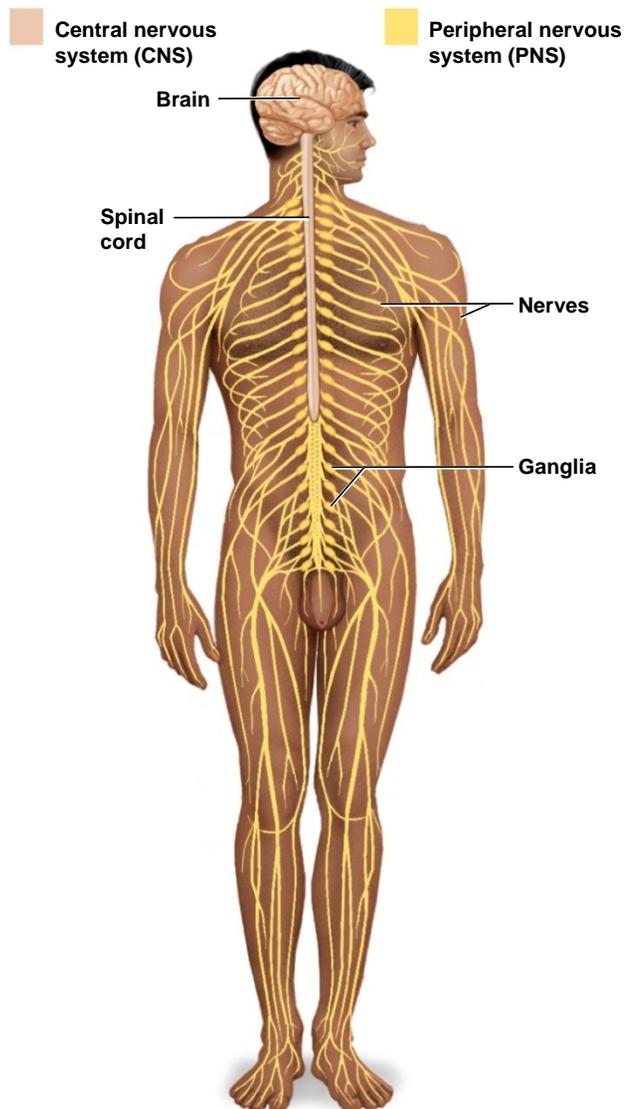
Overview of Nervous System

- endocrine and nervous system maintain internal coordination
 - **endocrine system** - communicates by means of chemical messengers (**hormones**) secreted into to the blood
 - **nervous system** - employs electrical and chemical means to send messages from cell to cell
 - nervous system carries out its task in **three basic steps**:
 - sense organs **receive information** about changes in the body and the external environment, and **transmits coded messages** to the spinal cord and the brain
 - brain and spinal cord **processes this information**, relates it to past experiences, and determine what response is appropriate to the circumstances
 - brain and spinal cord **issue commands** to muscles and gland cells to carry out such a response

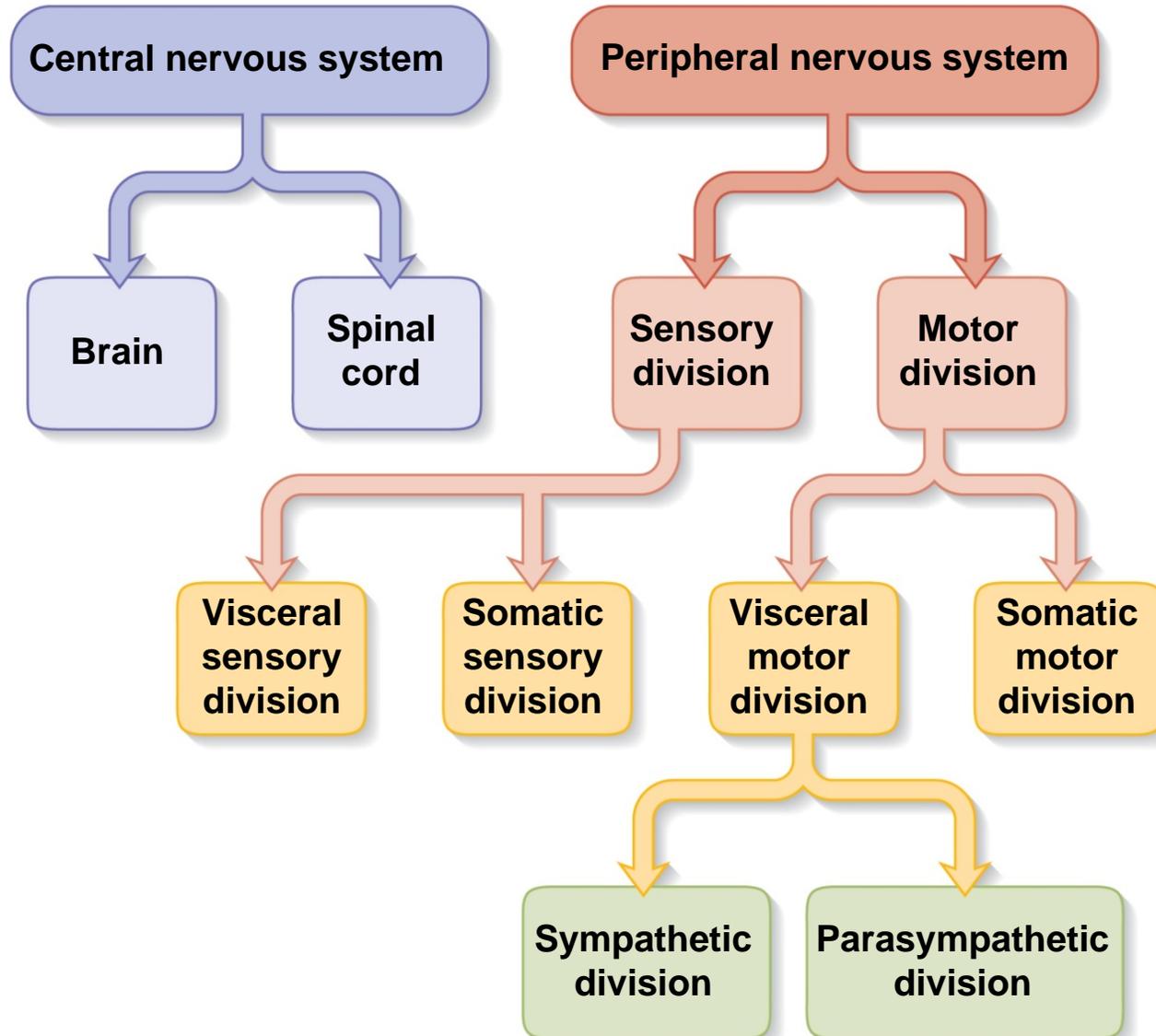
Two Major Anatomical Subdivisions of Nervous System

- **central nervous system (CNS)**
 - brain and spinal cord enclosed in bony coverings
 - enclosed by cranium and vertebral column
- **peripheral nervous system (PNS)**
 - all the nervous system except the brain and spinal cord
 - composed of **nerves** and **ganglia**
 - **nerve** – a bundle of nerve fibers (axons) wrapped in fibrous connective tissue
 - **ganglion** – a knot-like swelling in a nerve where neuron cell bodies are concentrated

Subdivisions of Nervous System



Subdivisions of Nervous System



Sensory Divisions of PNS

- **sensory division (afferent fibers)**
 - carries sensory signals from various receptors to the CNS
 - informs the CNS of stimuli within or around the body
 - **somatic sensory division** – carries signals from receptors in the skin, muscles, bones, and joints
 - **visceral sensory division** – carries signals from the viscera of the thoracic and abdominal cavities
 - heart, lungs, stomach, blood vessels, and urinary bladder

Motor Divisions of PNS

- **Motor division (efferent fibers)**
 - carries signals from the CNS to gland and muscle cells that carry out the body's response
 - **effectors** – cells and organs that respond to commands from the CNS
- **somatic motor division** – carries signals to skeletal muscles
 - output produces skeletal muscular contraction
 - as well as **somatic reflexes** – involuntary muscle contractions of skeletal muscles
- **visceral motor division (autonomic nervous system)**
 - carries signals to glands, cardiac muscle, and smooth muscle
 - involuntary, and responses of this system and its receptors are **visceral reflexes**
 - **sympathetic division**
 - tends to arouse body for action
 - accelerating heart beat and respiration, while inhibiting digestive and urinary systems
 - **parasympathetic division**
 - tends to have calming effect
 - slows heart rate and breathing
 - stimulates digestive and urinary systems

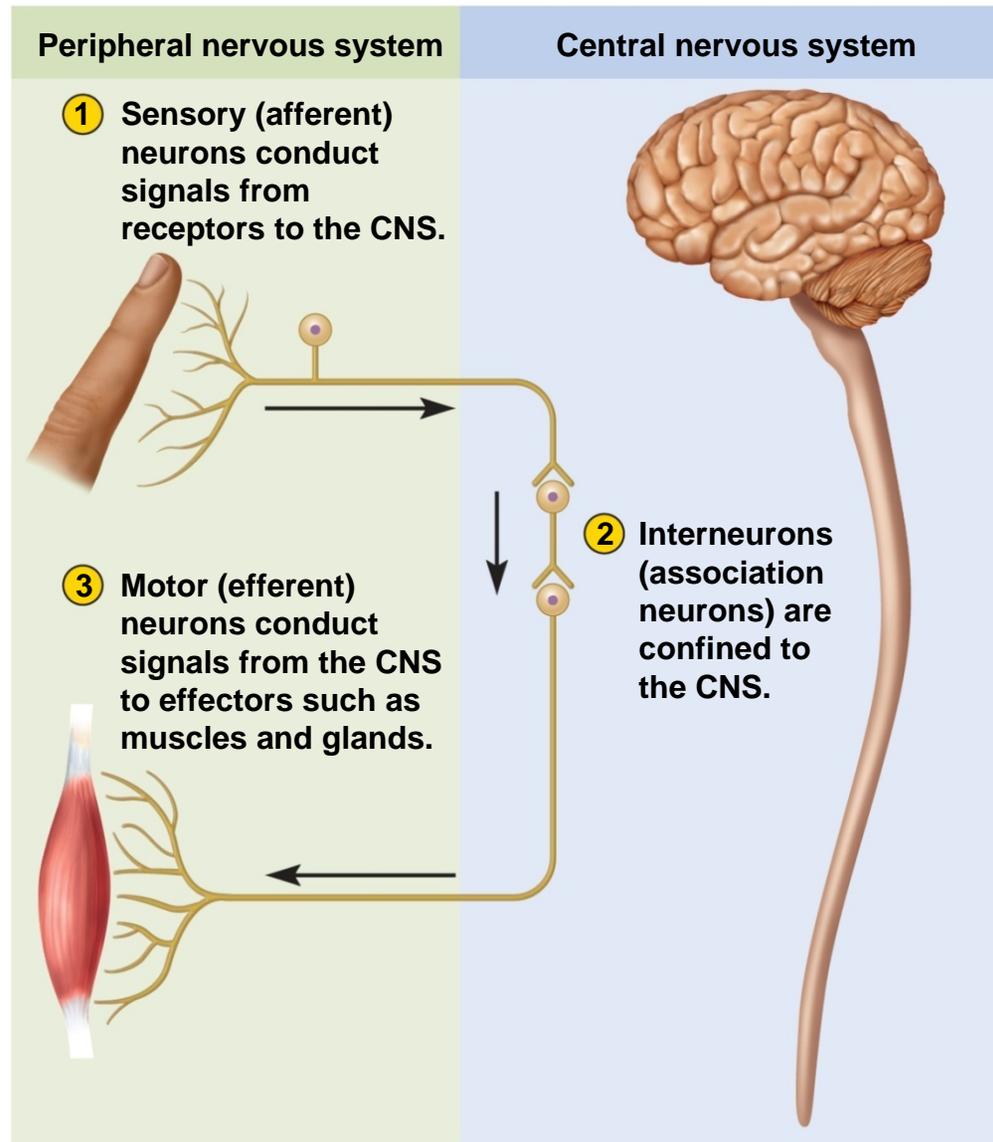
Universal Properties of Neurons

- **excitability (irritability)**
 - respond to environmental changes called **stimuli**
- **conductivity**
 - neurons respond to stimuli by producing **electrical signals** that are quickly conducted to other cells at distant locations
- **secretion**
 - when electrical signal reaches end of nerve fiber, a chemical **neurotransmitter** is secreted that crosses the gap and stimulates the next cell
- Note: electro-chemical communication!

Functional Types of Neurons

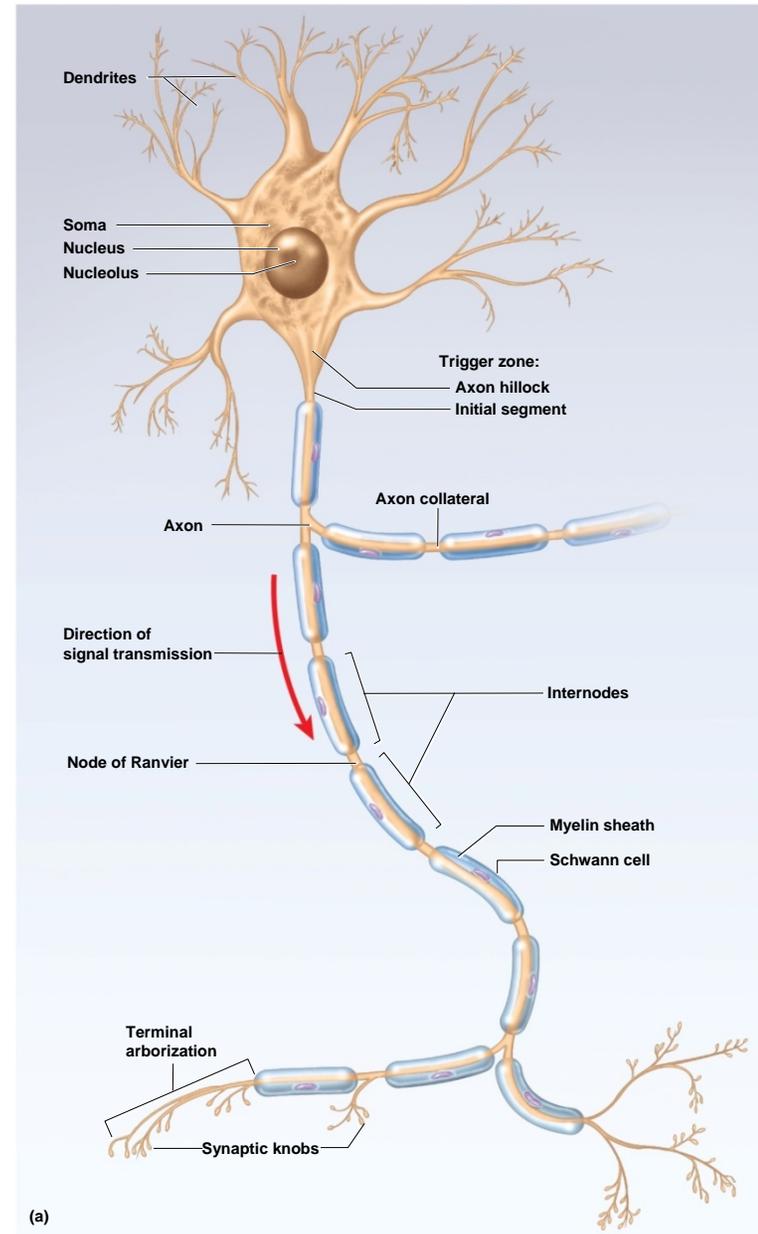
- **sensory (afferent) neurons**
 - specialized to detect stimuli
 - transmit information about them to the CNS
 - begin in almost every organ in the body and end in CNS
 - **afferent** – conducting signals toward CNS
- **interneurons (association) neurons**
 - lie entirely within the CNS
 - receive signals from many neurons and carry out the **integrative function**
 - process, store, and retrieve information and ‘make decisions’ that determine how the body will respond to stimuli
 - **90% of all neurons are interneurons**
 - lie between, and interconnect the incoming sensory pathways, and the outgoing motor pathways of the CNS
- **motor (efferent) neuron**
 - send signals out to muscles and gland cells (the **effectors**)
 - **motor** because most of them lead to muscles
 - **efferent** neurons conduct signals away from the CNS

Functional Classes of Neurons



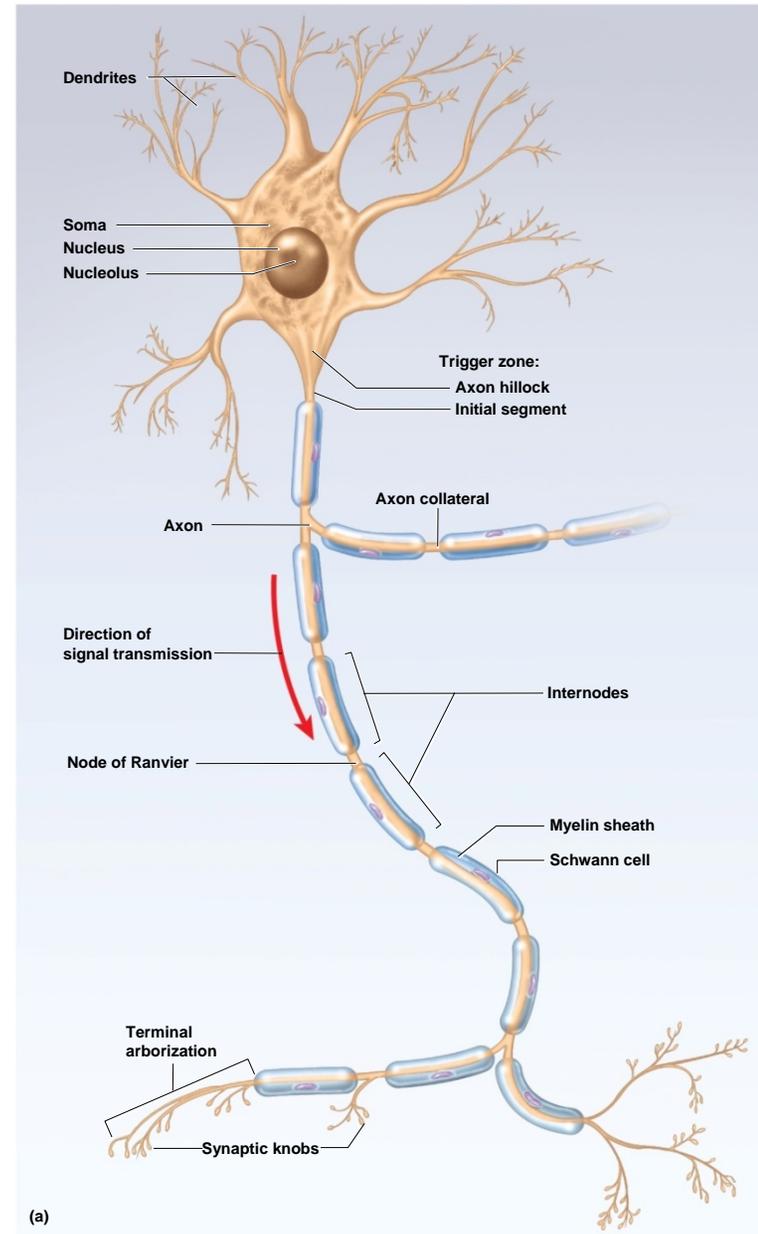
Structure of a Neuron – The Soma

- **soma** – the control center of the neuron
 - also called **neurosoma**, **cell body**, or **perikaryon**
 - has a single, centrally located nucleus with large nucleolus
 - **cytoplasm** contains mitochondria, lysosomes, a Golgi complex, numerous inclusions, and extensive rough endoplasmic reticulum and cytoskeleton
 - cytoskeleton consists of dense mesh of microtubules and **neurofibrils** (bundles of actin filaments)
 - compartmentalizes rough ER into dark staining **Nissl bodies**
 - no centrioles – no further cell division
 - **inclusions** – glycogen granules, lipid droplets, melanin, and **lipofuscin** (golden brown pigment produced when lysosomes digest worn-out organelles)
 - lipofuscin accumulates with age
 - wear-and-tear granules
 - most abundant in old neurons



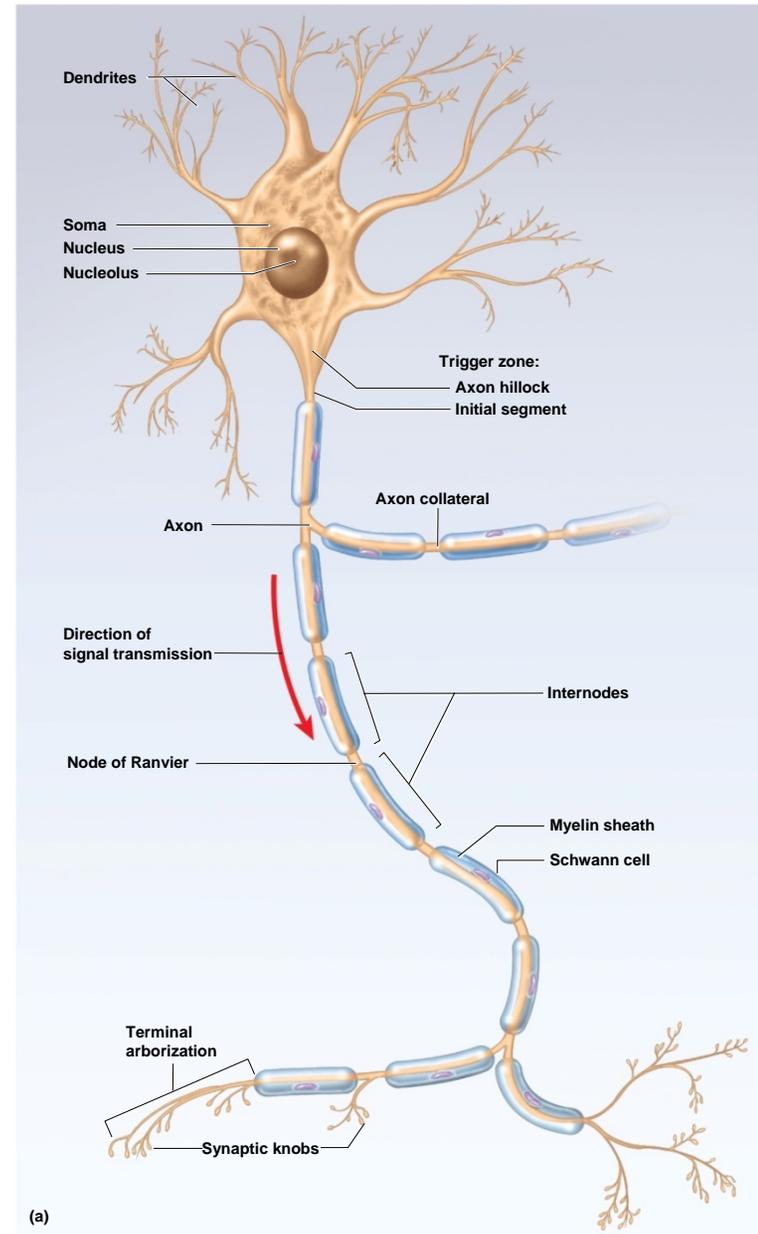
Structure of a Neuron – The Dendrites

- **dendrites** – vast number of branches coming from a few thick branches from the soma
 - resemble bare branches of a tree in winter
 - primary site for receiving signals from other neurons
 - the more dendrites the neuron has, the more information it can receive and incorporate into decision making
 - provide precise pathway for the reception and processing of neural information



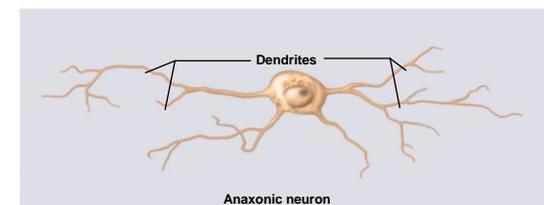
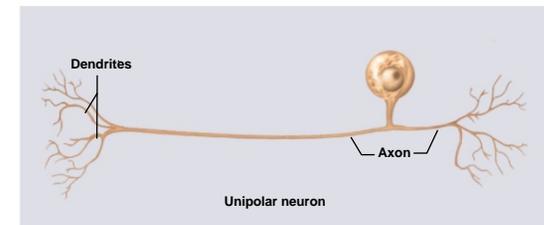
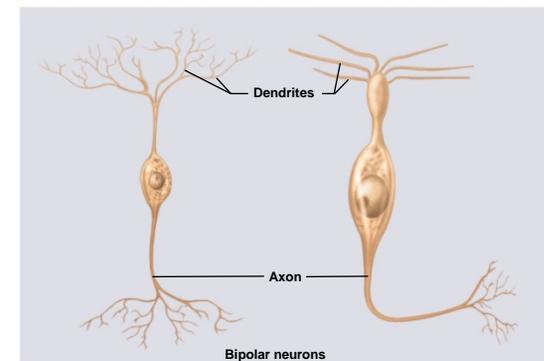
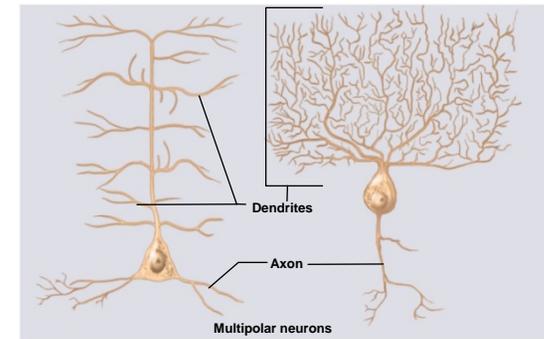
Structure of a Neuron – The Axon

- **axon (nerve fiber)** – originates from a mound on one side of the soma called the **axon hillock**
 - cylindrical, relatively unbranched for most of its length
 - **axon collaterals** – branches of axon
 - branch extensively on distal end
 - specialized for rapid conduction of nerve signals to points remote to the soma
 - **axoplasm** – cytoplasm of axon
 - **axolemma** – plasma membrane of axon
 - only **one axon per neuron**
 - **Schwann cells** and **myelin sheath** enclose axon
 - distal end, axon has **terminal arborization** – extensive complex of fine branches
 - **synaptic knob** (terminal button) – little swelling that forms a junction (synapse) with the next cell
 - contains **synaptic vesicles** full of neurotransmitter



Variation in Neuron Structure

- **multipolar neuron**
 - one axon and multiple dendrites
 - most common
 - most neurons in the brain and spinal cord
- **bipolar neuron**
 - one axon and one dendrite
 - olfactory cells, retina, inner ear
- **unipolar neuron**
 - single process
 - sensory from skin and organs to spinal cord
- **anaxonic neuron**
 - many dendrites but no axon
 - help in visual processes



Axonal Transport

- many **proteins made in soma** must be **transported to axon** and axon terminal
 - to repair axolemma, serve as gated ion channel proteins, as enzymes or neurotransmitters
- **axonal transport**
 - two-way passage of proteins, organelles, and other material along an axon
 - **anterograde transport** – movement down the axon away from soma
 - **retrograde transport** – movement up the axon toward the soma
- **microtubules** guide materials along axon
 - motor proteins (kinesin and dynein) carry materials “on their backs” while they **“crawl” along microtubules**
 - **kinesin** – motor proteins in anterograde transport
 - **dynein** – motor proteins in retrograde transport

Two Types of Axonal Transport

Fast and Slow

- **fast axonal transport**
 - occurs at a rate of 20 – 400 mm/day
 - **fast anterograde transport** (up to 400 mm/day)
 - organelles, enzymes, synaptic vesicles and small molecules
 - **fast retrograde transport**
 - for recycled materials and pathogens - rabies, herpes simplex, tetanus, polio viruses
 - delay between infection and symptoms is time needed for transport up the axon
- **slow axonal transport** or **axoplasmic flow**
 - 0.5 to 10 mm/day
 - always anterograde
 - moves enzymes, cytoskeletal components, and new axoplasm down the axon during repair and regeneration of damaged axons
 - damaged nerve fibers regenerate at a speed governed by slow axonal transport

Neuroglial Cells

- about a **trillion (10^{12}) neurons** in the nervous system (note: approximately 100 trillion in organism!)
- **neuroglia** outnumber the neurons by as much as **50 to 1**
- **neuroglia** or **glial cells**
 - support and protect the neurons
 - bind neurons together and form framework for nervous tissue
 - in fetus, guide migrating neurons to their destination
 - if mature neuron is not in synaptic contact with another neuron then it is covered by glial cells
 - prevents neurons from touching each other
 - gives precision to conduction pathways

Six Types of Neuroglial Cells (1 of 3)

- four types occur only in CNS
 - **oligodendrocytes**
 - form myelin sheaths in CNS
 - each arm-like process wraps around a nerve fiber forming an insulating layer that speeds up signal conduction
 - **ependymal cells**
 - lines internal cavities of the brain
 - cuboidal epithelium with cilia on apical surface
 - secretes and circulates cerebrospinal fluid (CSF)
 - clear liquid that bathes the CNS
 - **microglia**
 - small, wandering macrophages
 - formed by white blood cell called **monocytes**
 - thought to perform a complete checkup on the brain tissue several times a day
 - wander in search of cellular debris to phagocytize

Six Types of Neuroglial Cells (2 of 3)

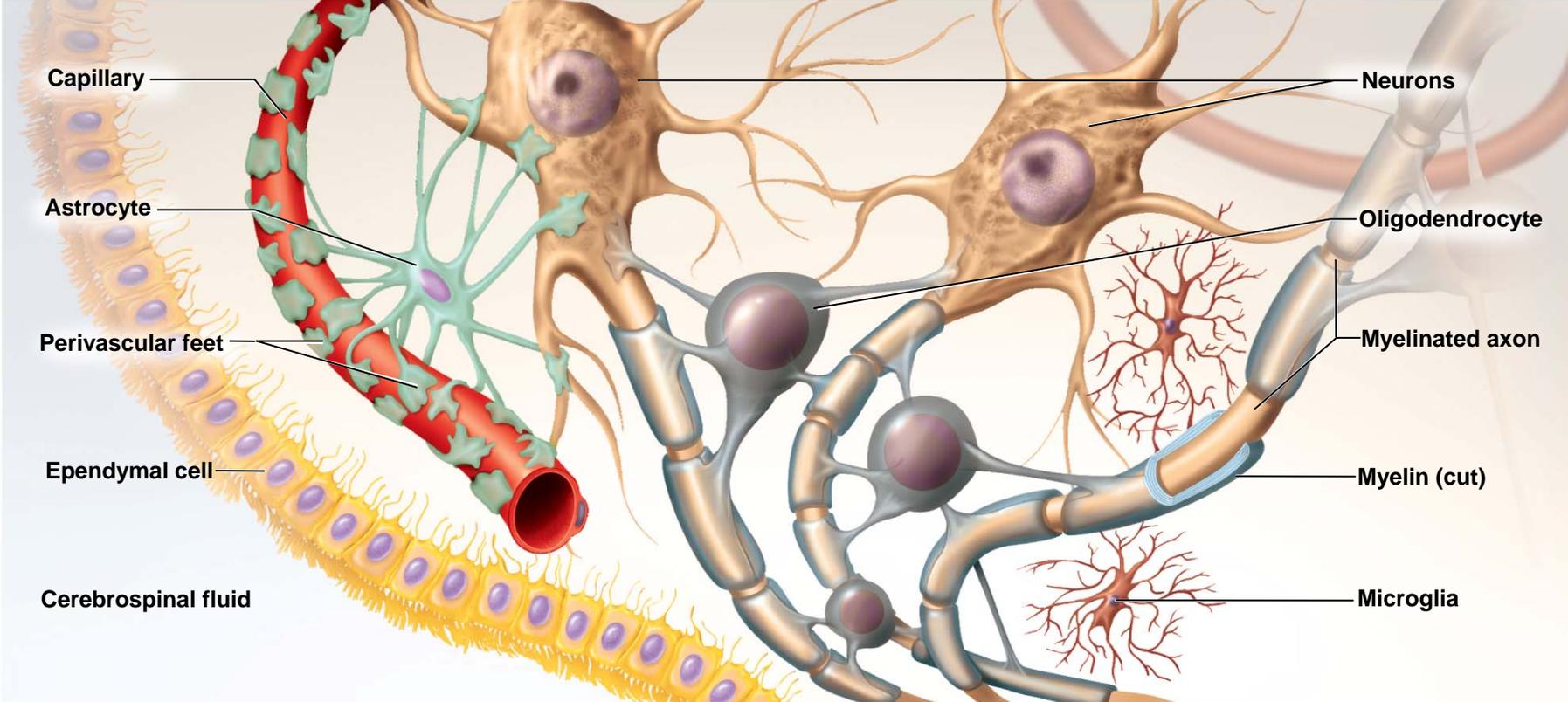
– astrocytes

- most abundant glial cell in CNS
- cover entire brain surface and most nonsynaptic regions of the neurons in the gray matter of the CNS
- diverse functions
 - form a supportive framework of nervous tissue
 - have extensions (**perivascular feet**) that contact blood capillaries that stimulate them to form a tight seal called the **blood-brain barrier**
 - convert blood glucose to **lactate** and supply this to the neurons for nourishment
 - **nerve growth factors** secreted by astrocytes promote neuron growth and synapse formation
 - communicate electrically with neurons and may influence synaptic signaling
 - regulate chemical composition of tissue fluid by absorbing excess neurotransmitters and ions
 - **astrocytosis** or **sclerosis** – when neuron is damaged, astrocytes form hardened scar tissue and fill space formerly occupied by the neuron

Six Types of Neuroglial Cells (3 of 3)

- two types occur only in PNS
 - **Schwann cells**
 - envelope nerve fibers in PNS
 - wind repeatedly around a nerve fiber
 - produces a **myelin sheath** similar to the ones produced by oligodendrocytes in CNS
 - assist in the regeneration of damaged fibers
 - **Satellite cells**
 - surround the neurosomas in ganglia of the PNS
 - provide electrical insulation around the soma
 - regulate the chemical environment of the neurons

Neuroglial Cells of CNS



Glial Cells and Brain Tumors

- **tumors** - masses of rapidly dividing cells
 - mature neurons have little or no capacity for mitosis and seldom form tumors
- **brain tumors** arise from:
 - meninges (protective membranes of CNS)
 - by metastasis from non-neuronal tumors in other organs
 - most come from glial cells that are mitotically active throughout life
- **gliomas** grow rapidly and are highly malignant
 - blood-brain barrier decreases effectiveness of chemotherapy
 - treatment consists of radiation or surgery

Myelin

- **myelin sheath**
 - an **insulating layer** around a nerve fiber
 - formed by **oligodendrocytes in CNS** and **Schwann cells in PNS**
 - consists of the **plasma membrane of glial cells**
 - 20% protein and 80 % lipid
- **Myelination**
 - production of the myelin sheath
 - begins the **14th week of fetal development**
 - proceeds rapidly during infancy
 - completed in late adolescence
 - dietary fat is important to nervous system development (Danger! Trans Fat!)

Myelin

- In PNS
 - **Schwann cell** spirals repeatedly around a single nerve fiber
 - lays down as many as a hundred layers of its own membrane
 - no cytoplasm between the membranes
 - **neurilemma** – thick outermost coil of myelin sheath
 - contains nucleus and most of its cytoplasm
 - external to neurilemma is basal lamina and a thin layer of fibrous connective tissue – **endoneurium**

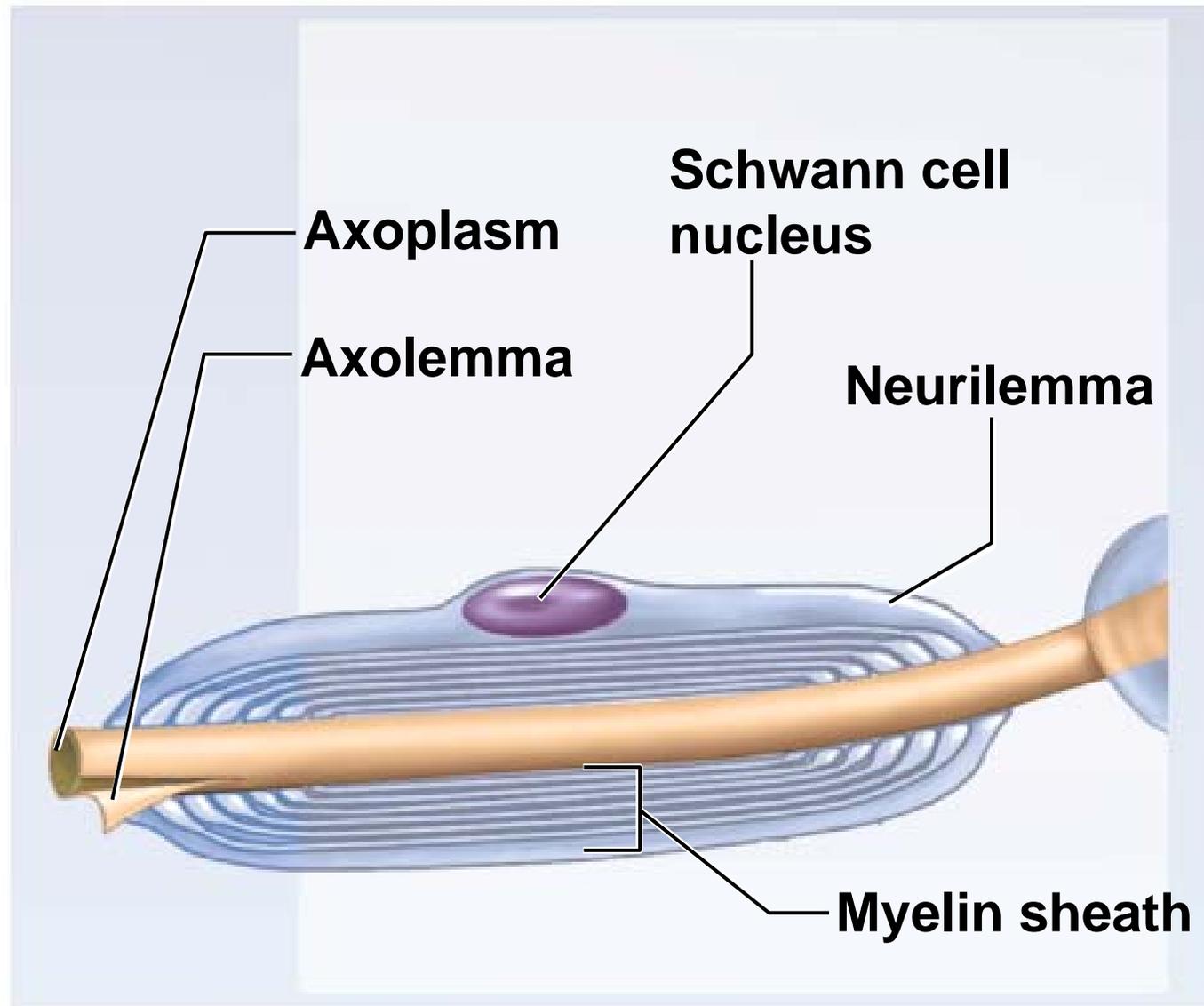
Myelin

- In CNS
 - **oligodendrocytes** reaches out to myelinate several nerve fibers in its immediate vicinity
 - anchored to multiple nerve fibers
 - cannot migrate around any one of them like Schwann cells
 - must push newer layers of myelin under the older ones
 - so myelination spirals inward toward nerve fiber
 - nerve fibers in CNS have **no neurilemma or endoneurium**

Myelin

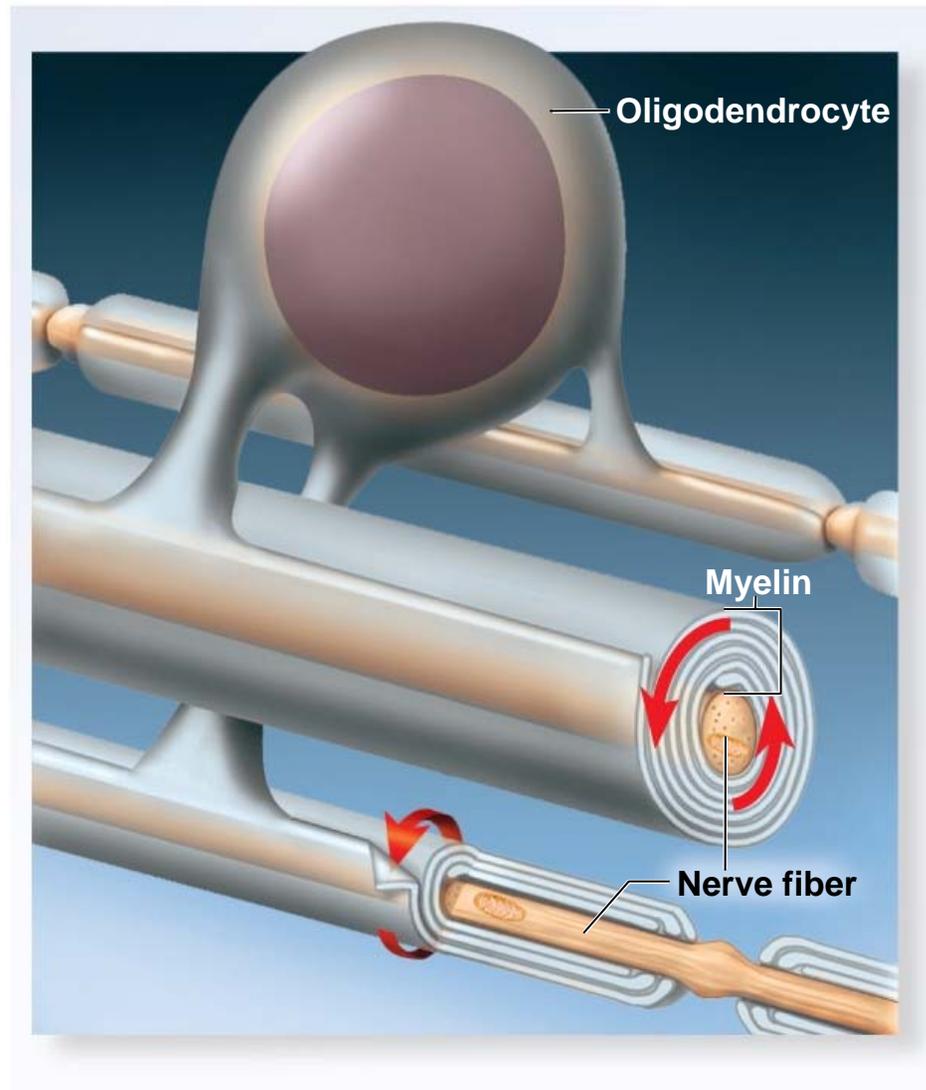
- Many Schwann cells or oligodendrocytes are needed to cover one nerve fiber
 - Schwann cells in PNS
 - Oligodendrocytes in CNS
- **myelin sheath is segmented**
 - **nodes of Ranvier** – gap between segments
 - **internodes** – myelin covered segments from one gap to the next
 - **initial segment** – short section of nerve fiber between the axon hillock and the first glial cell
 - **trigger zone** – the axon hillock and the initial segment
 - play an important role in initiating a nerve signal

Myelin Sheath in PNS



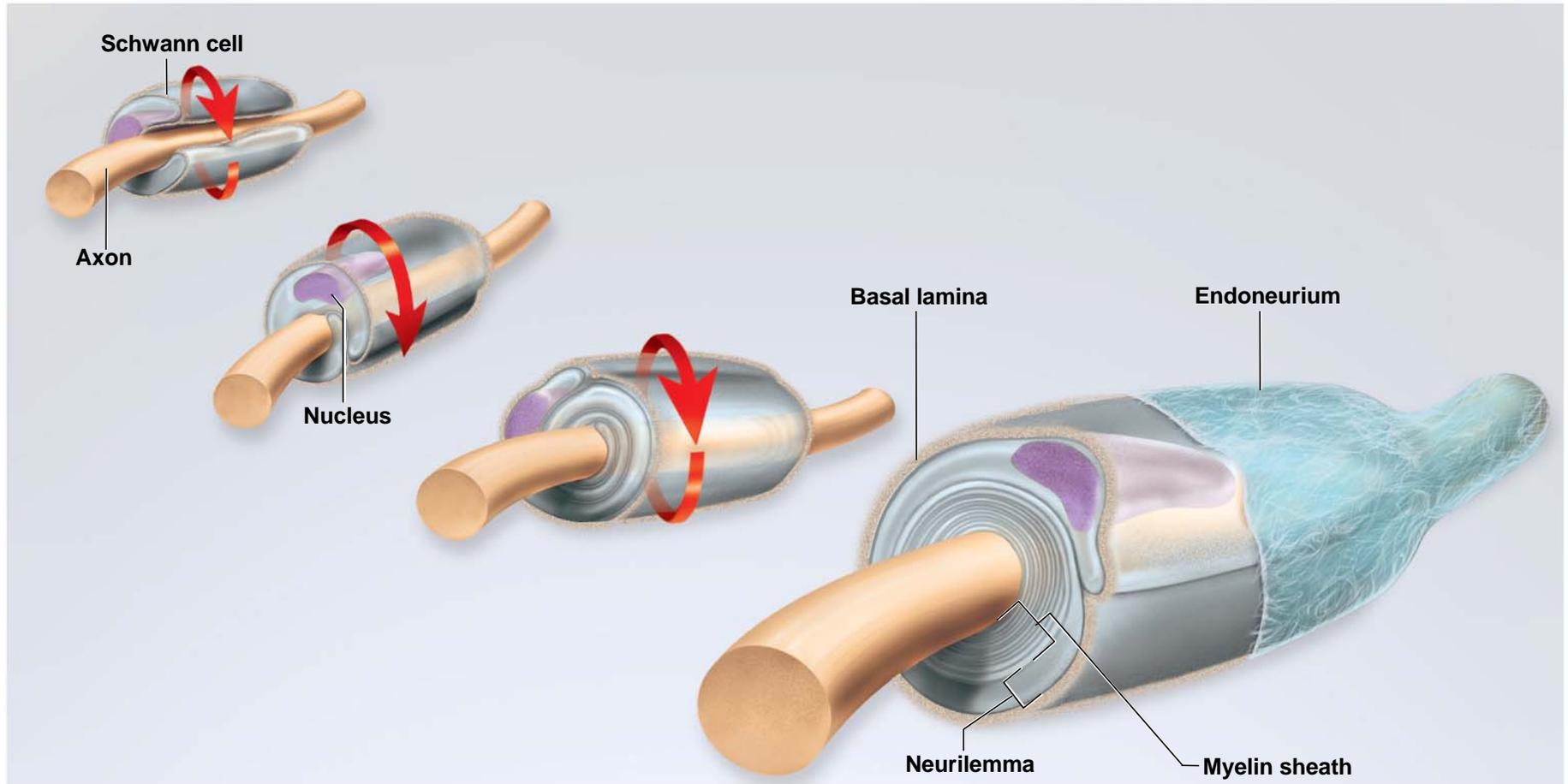
nodes of Ranvier and internodes

Myelination in CNS



Myelination in PNS

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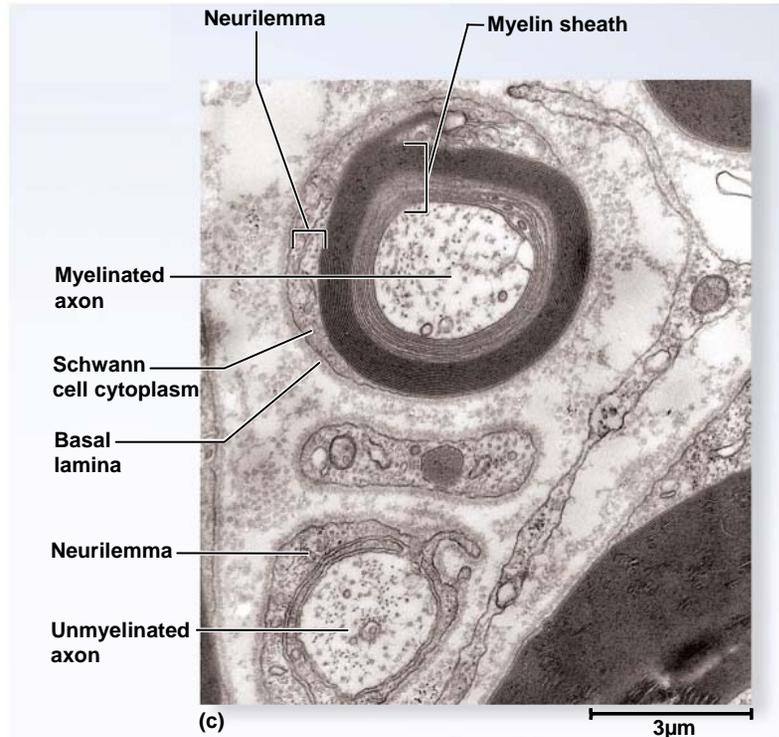


Diseases of Myelin Sheath

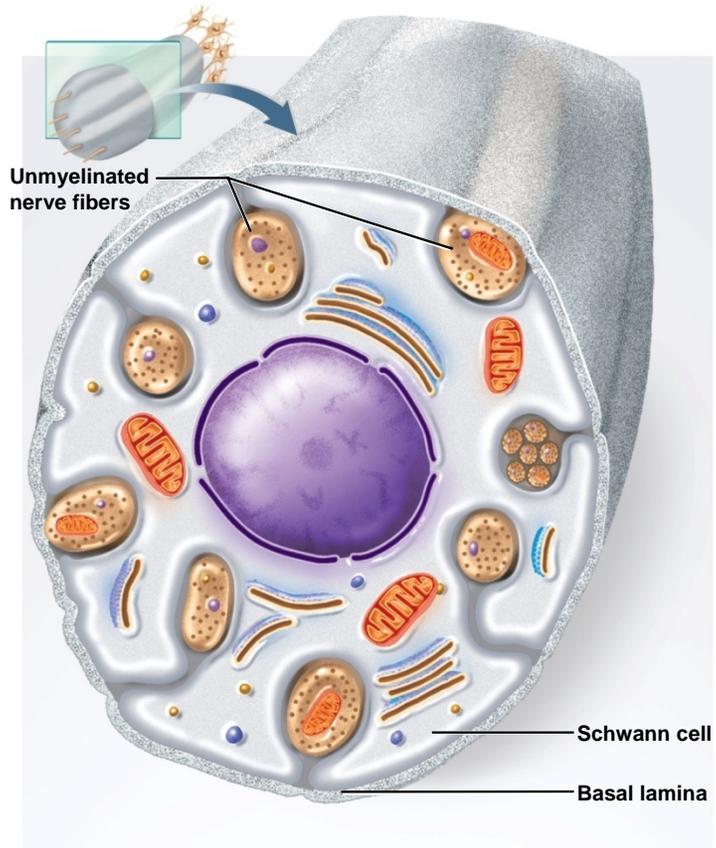
- degenerative disorders of the myelin sheath
 - **multiple sclerosis**
 - oligodendrocytes and myelin sheaths in the CNS deteriorate
 - myelin replaced by hardened scar tissue
 - nerve conduction disrupted (double vision, tremors, numbness, speech defects)
 - onset between 20 and 40 and fatal from 25 to 30 years after diagnosis
 - cause may be autoimmune triggered by virus
 - **Tay-Sachs disease** - a hereditary disorder of infants of Eastern European Jewish ancestry
 - abnormal accumulation of glycolipid called **GM₂** in the myelin sheath
 - normally decomposed by lysosomal enzyme
 - enzyme missing in individuals homozygous for Tay-Sachs allele
 - accumulation of **ganglioside** (GM₂) disrupts conduction of nerve signals
 - blindness, loss of coordination, and dementia
 - fatal before age 4

Unmyelinated Axons of PNS

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- Schwann cells hold 1 – 12 small nerve fibers in grooves on its surface
- membrane folds once around each fiber overlapping itself along the edges
- **mesaxon** – neurilemma wrapping of unmyelinated nerve fibers

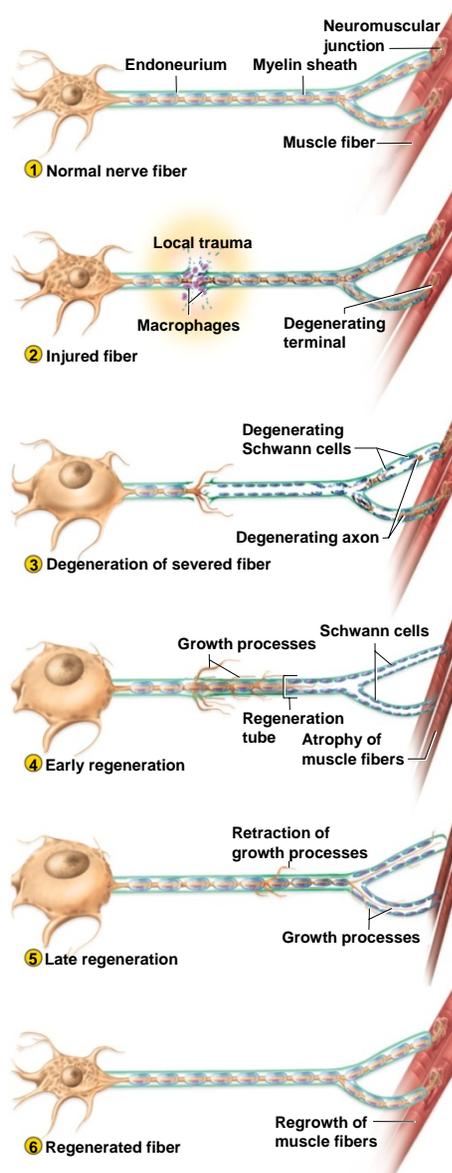
Conduction Speed of Nerve Fibers

- speed at which a nerve signal travels along a nerve fiber depends on two factors
 - **diameter of fiber**
 - **presence or absence of myelin**
- signal conduction occurs along the surface of a fiber
 - larger fibers have more surface area and conduct signals more rapidly
 - myelin further speeds signal conduction
- **conduction speed**
 - small, unmyelinated fibers - 0.5 - 2.0 m/sec
 - small, myelinated fibers - 3 - 15.0 m/sec
 - large, myelinated fibers - up to 120 m/sec
 - slow signals supply the stomach and dilate pupil where speed is less of an issue
 - fast signals supply skeletal muscles and transport sensory signals for vision and balance

Regeneration of Peripheral Nerves

- **regeneration** of a damaged **peripheral nerve fiber** can occur if:
 - its **soma is intact**
 - at least **some neurilemma remains**
- fiber distal to the injury cannot survive and it degenerates
 - macrophages clean up tissue debris at the point of injury and beyond
- soma swells, ER breaks up, and nucleus moves off center
 - **due to loss of nerve growth factor from neuron's target cell**
- axon stump sprouts multiple growth processes
 - severed distal end continues to degenerate
- **regeneration tube** – formed by Schwann cells, basal lamina, and the neurilemma near the injury
 - **regeneration tube guides the growing sprout back to the original target cells and reestablishes synaptic contact**
- nucleus returns to normal shape
- Note: regeneration of damaged nerve fibers in the **CNS cannot occur at all**

Regeneration of Nerve Fiber



denervation atrophy

Atrophy of muscle due to loss of nerve contact by damaged nerve

Electrophysiology of Neurons

- **Galen** thought that the brain pumped a vapor called *psychic pneuma* through hollow nerves and squirted in to the muscles to make them contract
- **Rene' Descartes** in the 17th century supported this theory
- **Luigi Galvani** discovered the role of electricity in muscle contraction in the 18th century
- **Camillo Golgi** developed important method for staining neurons with silver in the 19th century
- **Santiago Ramon y Cajal** set forth the **neuron doctrine** – nervous pathway is not a continuous 'wire' or tube, but a series of cells separated by gaps called synapses.
- neuron doctrine brought up two key questions:
 - how does a neuron generate a electrical signal?
 - how does it transmit a meaningful message to the next cell?

Nerve Growth Factor

- **nerve growth factor (NGF)**
 - a **protein secreted by a gland, muscle, and glial cells** and picked up by the axon terminals of the neurons.
 - **prevents apoptosis** (programmed cell death) in growing neurons
 - enables growing neurons to make contact with their target cells
- isolated by Rita Levi-Montalcini in 1950s
- won Nobel prize in 1986 with Stanley Cohen
- use of growth factors is now a vibrant field of research