# Cells

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- Treat the tissue with a preservative (e.g. formaldehyde).
- Dissect the region of interest.
- Embed the tissue.
- Cut the tissue into thin sections.
- Stain the tissue to reveal subject of interest.
- Examine with a microscope.

 Different stains show different cell types, cell organelles or molecules.



 This purple stain shows myelin, which is on axons.

This purple stain shows neuronal cell bodies.



## Microscopy



5

specimen

## Microscopy

• electron microscope





#### Cells



- Cells are small – cell bodies (somas) of neurons range from 10 to  $100 \mu m$  in diameter

- A cell has organelles such as a nucleus and mitochondria (as a person has organs such as a brain and a liver)
- The organelles of the cells of the nervous system are like those of most other cells of the body.

Cell membrane (plasma membrane):

- Bounds the entire surface of the cell including the axon and dendrites of neurons.
- Its integrity is essential for normal cell function.



Cell membrane:

- Composed of a phospholipid bilayer, which forms an impermeable barrier to water and ions.
- Many proteins pass through or are attached to the membrane including:
  - Cell adhesion molecules
  - Ion channels & pumps
  - Receptors







Nucleus:

• The most conspicuous organelle of a cell.



Nucleus includes:

- Nuclear envelope with pores, which surrounds the nucleus and allows selected molecules to pass between the nucleus and cytoplasm
- Nucleolus, which has important roles in genesis of ribosomes
- Chromosomes, which are double strands of DNA complexed with proteins



Nucleus:

Low magnification light micrograph



Medium magnification Molecular layer light micrograph PC Electron micrograph Granule cells Primary dendrite Purkinje cell Granule cells

5 µm

### Nucleus:

Low magnification fluorescence micrograph



Chromosomes:

- Humans normally have 23 pairs of chromosomes, the last of which are the sex chromosomes.
- Females are XX, and males are XY.
- Deletion or duplication of a chromosome can cause abnormalities.

A third copy of chromosome 21 results in Down's syndrome, a common cause of mental retardation.



**DNA & Genes** 

- DNA is a helical chain of four types of nucleotides in a specific sequence.
- Many segments of the DNA of a chromosome are genes. A gene encodes the instructions for synthesis of a protein.
- Humans have ~23,000 genes.
- Genes are the basis of heredity.
- Most cells of an individual have the same DNA (genes).



Protein synthesis:

- A gene is used as a template for synthesis of messenger RNA (mRNA) in the nucleus
- mRNA is used as a template for synthesis of a protein in the cytoplasm.

DNA (gene) > mRNA > protein

• Different cell types express only certain proteins. The function of a cell is determined by the proteins it expresses.



- DNA is a chain of four nucleotides.
- mRNA is a chain of four slightly different nucleotides.
- Protein is a chain of amino acids.
- The sequence of three nucleotides in the mRNA specifies the amino acid to assemble into the protein being synthesized.
- Thus, the sequence of nucleotides in a gene (DNA) ultimately determines the sequence of amino acids in a protein.



- Synthesis of mRNA from DNA in the nucleus is a process called <u>transcription</u>.
- Synthesis of protein from mRNA in the cytoplasm is a process called <u>translation</u>.



Ribosome:

• Reads the sequence of mRNA to synthesize proteins.



Endoplasmic reticulum:

- Interconnected network of tubules in the cytoplasm
- Rough endoplasmic reticulum have ribosomes on its surface and is important for synthesis of membrane proteins and proteins for export.



- Neurons have large amounts of rough endoplasmic reticulum in their somas, which we call Nissl substance.
- Nissl substance is readily seen by microscopy.
- Most proteins and other molecules needed by neurons are synthesized in the soma.



NB

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#### **Organelles of Cells**

Nissl stain:



Low magnification light micrograph of spinal cord High magnification light micrograph of a spinal neuron



Golgi apparatus:

- Flattened membrane bound sacs
- Modifies newly synthesized proteins and lipids
- Gives rise to vesicles for transport throughout the cell



Mitochondria:

• Supply energy in a form that can be used by most enzymes in the cell



Lysosome:

• Breaks down organelles and molecules for recycling.



Cytoskeleton:

- Filaments of structural proteins present in a matrix throughout the cell.
- Maintains the shape of the cell, involved in the transport of molecules and organelles, and involved in cell movement.
- Three main elements:
  - Microtubules (tubulin)
  - Intermediate filaments (neurofilament protein in neurons)
  - Microfilaments (actin)



## microtubules nucleus



- dendrite
- soma
- axon
- synapse

- In neurons, most protein synthesis takes place in the cell body.
- Very little mRNA is present in the axon.



- Yet most of the volume of a neuron can be in its axon.
- Axonal transport is required to move molecules and organelles around the cell.





• Different components of the cytoskeleton are involved in transport within the axon to and from the soma.

Rabies:

- Rabies is a virus usually introduced through the skin via an animal bite.
- The virus is taken up by peripheral nerves and taken by axonal transport into the central nervous system where it has its primary effect.
- When the virus reaches the soma of a neuron, it releases its mRNA, which leads to synthesis of viral proteins.
- The viral proteins damage the host neuron and lead to production and release of more viruses.



Herpes simplex:

- A virus responsible for genital herpes, a sexually transmitted disease (STD).
- It is transmitted between partners during unprotected sex.
- It is transported in sensory axons, and will survive in the soma of sensory neurons for life.



Synapse or synaptic terminal

- Termination of an axon
- Site of communication with another cell (The presynaptic cell communicates with the postsynaptic cell.)
- Postsynaptic cell can be another neuron or other cell type.

- Synapses between neurons can be:
  - axodendritic
  - axosomatic
  - axoaxonic
  - dendrodendritic



synapse or

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Microtubules

synaptic terminal

Structure of a typical synapse:

- Presynaptic terminal
  - Synaptic vesicles containing neurotransmitter
  - Presynaptic density
- Synaptic cleft
- Postsynaptic element
  - Neurotransmitter receptors
  - Postsynaptic density

Synantic vesicles

Tubules of smooth ER

Neurofilaments



**Action Potential** 

• When the neuron is activated, an electrical pulse, the action potential, is carried down the axon.

Synaptic Communication

- When an action potential reaches a synapse, synaptic vesicles fuse with the cell membrane and release neurotransmitter into the synaptic cleft.
- Neurotransmitter crosses the synaptic cleft and interacts with receptors on the postsynaptic cell.
- Neurotransmitter will activate or inhibit the postsynaptic cell.

Different     neurotrans			neurons	release	different	
<ul> <li>Some common neurotransmitters:</li> </ul>						
class	transmitter					
biogenic amines acetylcholine						
	dopamine					
		norepinephrine (noradrenaline)				
		epinephrine (adrenaline)				
	:	serotonin				
amino acio	ls '	γ-aminobutyric acid (GABA)				
		glutamate				
		glyd	cine			
peptides	,	vasoactive intestinal polypeptide substance P				
	:					
		enkephalin				
	endorphin					

- Neurochemical communication requires the postsynaptic terminal to have the proper receptor for the neurotransmitter.
- The transmitter-receptor pair determines whether the active synapse will excite or inhibit the postsynaptic cell.



## Glia (Neuroglia) (The Non-neuronal Cells of the Nervous System)

- Central nervous system (CNS) includes the brain, spinal cord and retina.
- Peripheral nervous system (PNS) includes nerves and ganglia, which are distributed throughout the body.



## Glia (Neuroglia) (The Non-neuronal Cells of the Nervous System)

CNS glia:

- Astrocytes support the neurons
- Oligodendrocytes myelinate axons
- Ependymal cells line the ventricular system
- Microglia housekeepers

PNS glia:

- Satelite cells support the neurons
- Schwann cells myelinate axons
- Macrophages housekeepers

## Astrocytes

- Star-shaped glial cells in the CNS
- Most abundant cell type of the brain and spinal cord
- Surround most neurons and blood vessels
- Contribute to the cellular scaffolding of the CNS



• Mediate exchange between capillaries and neurons; contribute to the blood-brain barrier.



#### Astrocytes

