Modern Cell Theory

- All organisms composed of cells and cell products.
- Cell is the simplest structural and functional unit of life.
 cells are alive
- An organism's structure and functions are due to the activities of its cells.
- Cells come only from preexisting cells, not from nonliving matter.
 - therefore, all life traces its ancestry to the same original cells
- Cells of all species have many fundamental similarities in their chemical composition and metabolic mechanisms.

Cell Shapes

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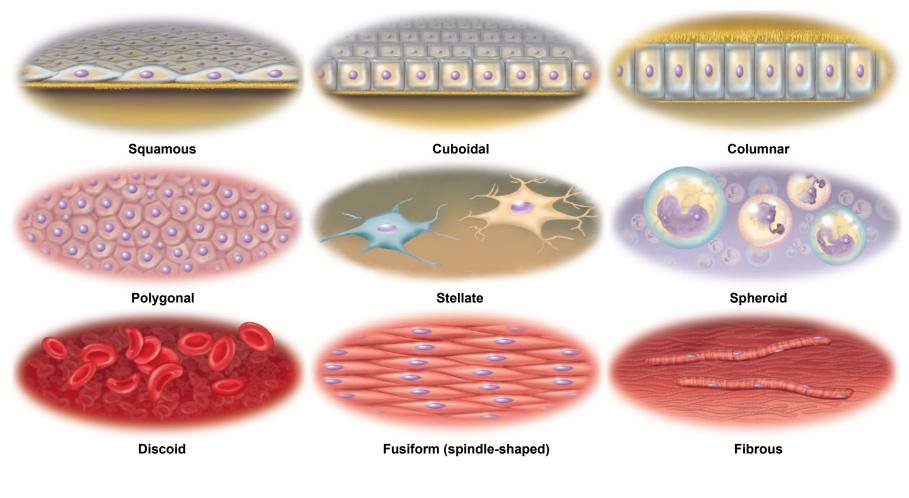


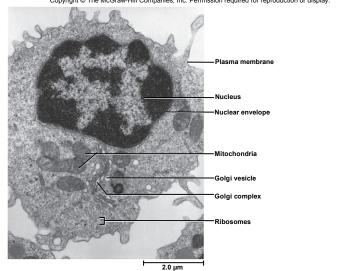
Figure 3.1

Cell Size

- Human cell size
 - 10 15 micrometers (μ m) in diameter
 - egg cells (very large)100 µm diameter
 - barely visible to the naked eye
 - nerve cell at 1 meter long
- Limitations on cell size
 - cell growth increases volume more than surface area
 - nutrient absorption and waste removal utilize surface area

General Cell Structure

- Light microscope reveals plasma membrane, nucleus and cytoplasm
 - cytoplasm fluid between the nucleus and surface membrane
- **Resolution** (ability to reveal detail) of electron microscopes reveals **ultrastructure**
 - organelles, cytoskeleton and cytosol (ICF)

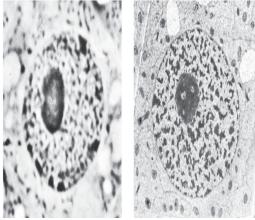


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Figure 3.3

Magnification versus Resolution

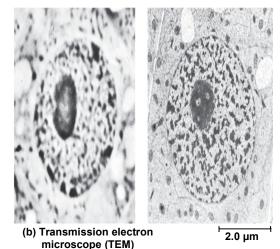
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(a) Light microscope (LM)
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Figure 3.4a

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Figure 3.4b

- **Magnification** of 750x seen through both light and transmission electron microscope
- Increased resolution reveals the finer details

Major Constituents of Cell

plasma (cell) membrane

- surrounds cell
- made of proteins and lipids
- composition and function can vary from one region of the cell to another

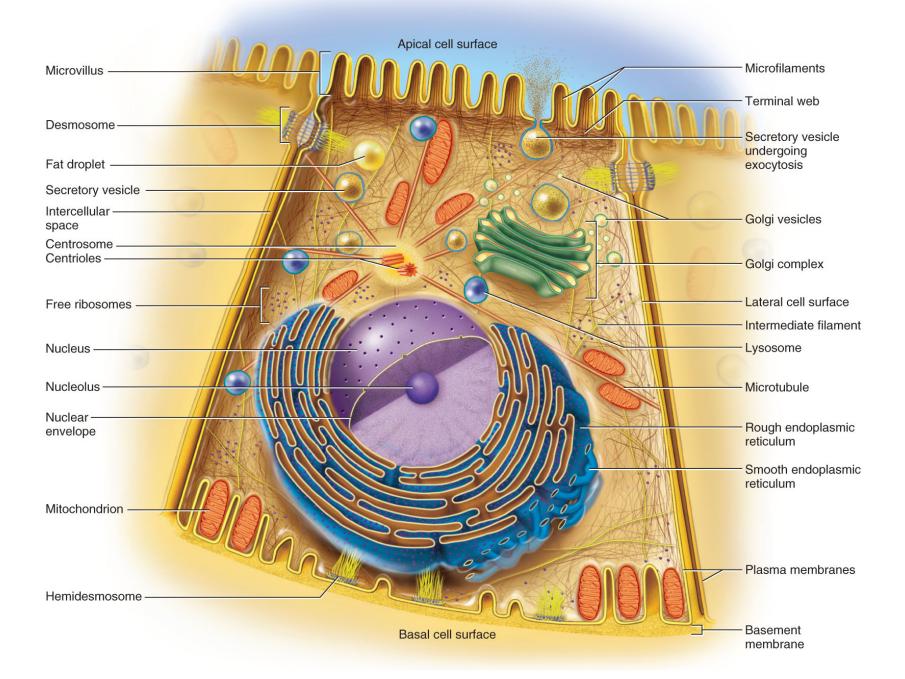
cytoplasm

- organelles
- cytoskeleton
- cytosol (intracellular fluid ICF)

extracellular fluid – ECF

- fluid outside of cell

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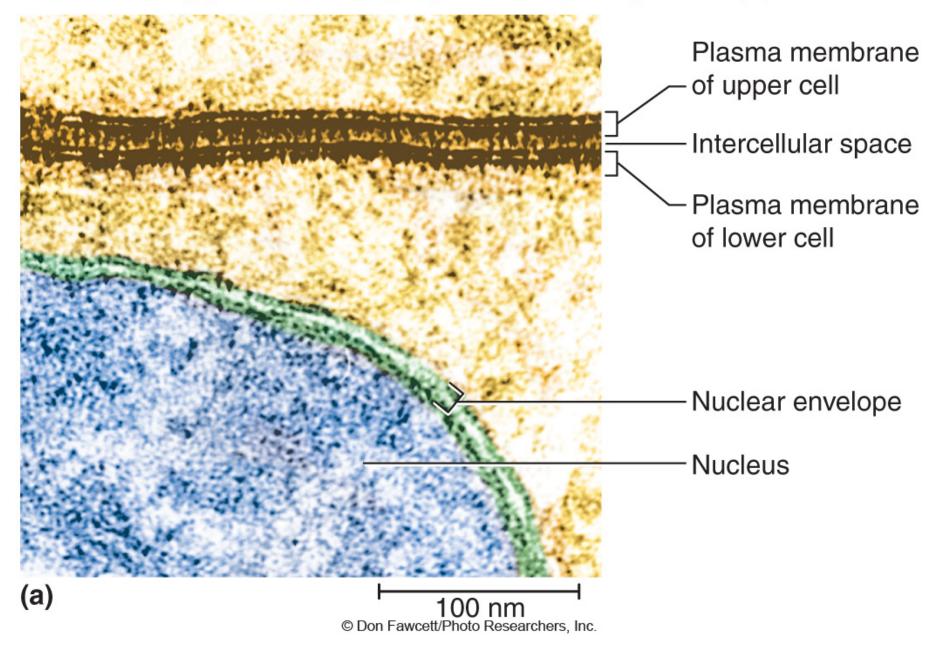


Plasma Membrane

plasma membrane – membrane at cell surface

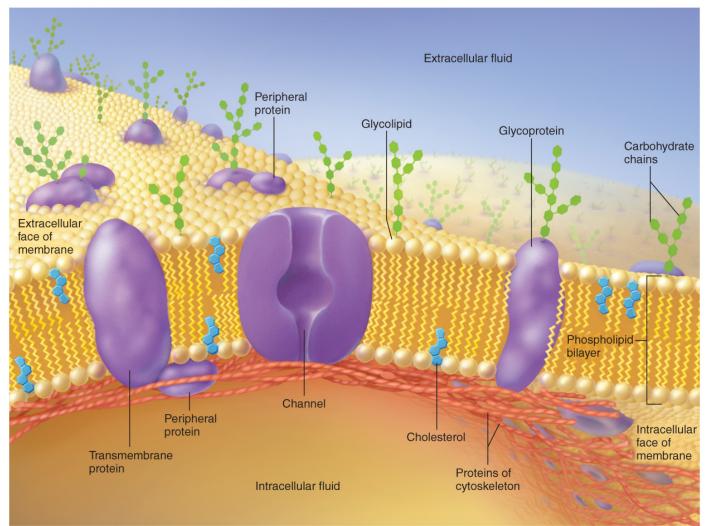
- -defines cell boundaries
- -governs interactions with other cells
- -controls passage of materials in and out of cell
- -intracellular face side that faces cytoplasm
- -extracellular face side that faces outward

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Plasma Membrane

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Oily film of lipids with diverse proteins embedded 3-10

Membrane Lipids

Phospholipids

- 75% of membrane lipids are phospholipids
- amphiphilic molecules arranged in a bilayer
 - hydrophilic phosphate heads face water on each side of membrane
 - hydrophobic tails directed toward the center, avoiding water
- drift laterally from place to place
- movement keeps membrane fluid

Cholesterol

- 20% of the membrane lipids
- holds phospholipids still and can stiffen membrane

Glycolipids

- 5% of the membrane lipids
- phospholipids with short carbohydrate chains on extracellular face
- contributes to glycocalyx carbohydrate coating on the cells surface

Membrane Proteins

Membrane proteins

- 2% of the molecules in plasma membrane
- 50% of its weight

Transmembrane proteins

- pass through membrane
- have hydrophilic regions in contact with cytoplasm and extracellular fluid
- have hydrophobic regions that pass back and forth through the lipid of the membrane
- most are glycoproteins
- can drift about freely in phospholipid film
- some anchored to cytoskeleton
- Peripheral proteins
 - adhere to one face of the membrane
 - usually tethered to the cytoskeleton

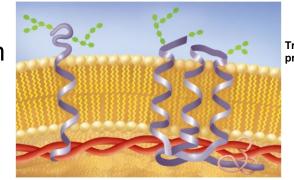
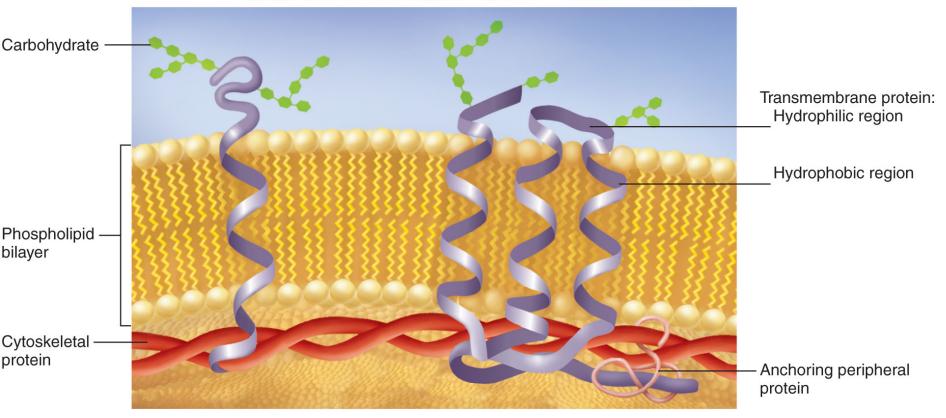


Figure 3.7 3-12

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Transmembrane protein:

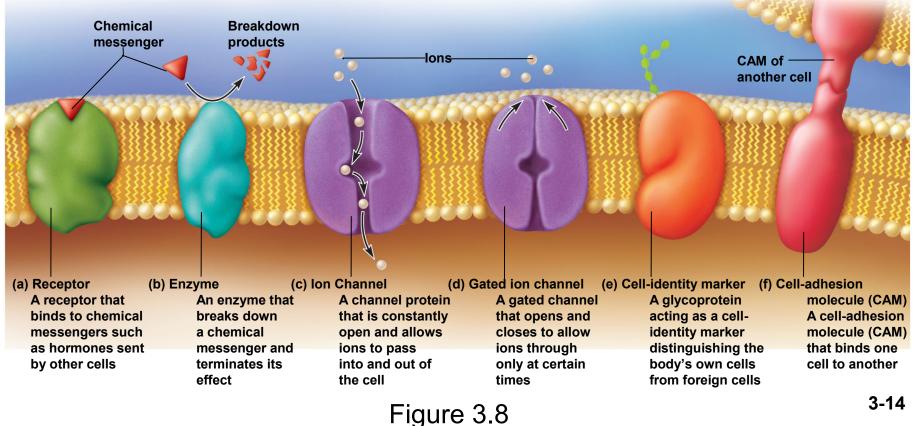


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Membrane Protein Functions

 receptors, second-messenger systems, enzymes, ion channels, carriers, cell-identity markers, cell-adhesion molecules

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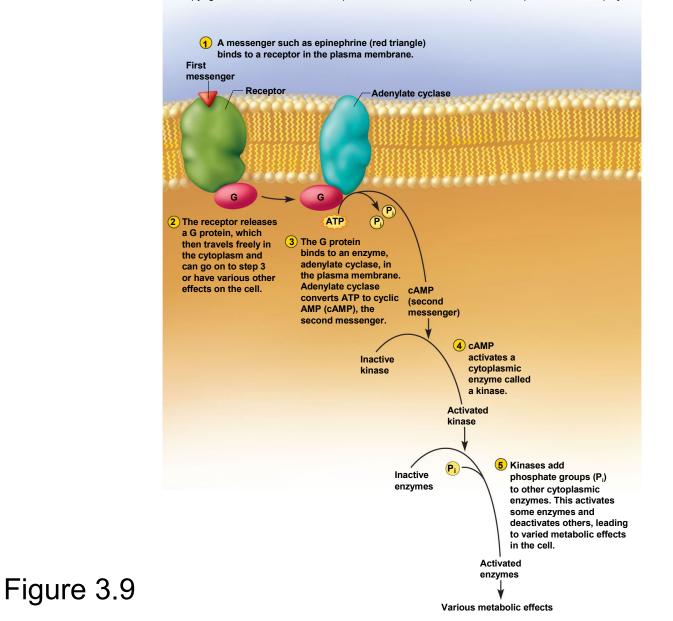


Membrane Receptors

- Cell communication via chemical signals
 - receptors surface proteins on plasma membrane of target cell
 - bind these chemicals (hormones, neurotransmitters)
 - receptor usually specific for one substrate

Second Messenger System

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3-16

Second Messenger System

- Chemical **first messenger** (epinephrine) binds to a surface receptor
 - triggers changes within the cell that produces a second messenger in the cytoplasm
- **Receptor** activates **G protein**
 - an intracellular peripheral protein
 - guanosine triphosphate (GTP) an ATP like compound
- G protein relays signal to adenylate cyclase which converts ATP to cAMP (2nd messenger)
- cAMP activates a kinase in the cytosol
- Kinases add phosphate groups to other cellular enzymes
 - activates some enzymes, and inactivates others triggering a wide variety of physiological changes in cells
- Up to 60% of modern drugs work by altering activity of G proteins.

Membrane Enzymes

- enzymes in plasma membrane carry out final stages of starch and protein digestion in small intestine
- help produce second messengers (cAMP)
- break down chemical messengers and hormones whose job is done
 - stops excessive stimulation

Ion Channels

- Transmembrane proteins with pores that allow water and dissolved ions to pass through membrane
 - some constantly open
 - some are gated-channels that open and close in response to stimuli
 - **ligand** (chemically)-regulated gates
 - voltage-regulated gates
 - **mechanically** regulated gates (stretch and pressure)
- play an important role in the timing of nerve signals and muscle contraction

Membrane Carriers or Pumps

• Transmembrane proteins bind to glucose, electrolytes, and other solutes

- transfer them across membrane

• Pumps consume ATP in the process

Cell-Identity Markers

- Glycoproteins contribute to the glycocalyx
 carbohydrate surface coating
 - acts like a cell's 'identification tag'
- Enables our bodies to identify which cells belong to it and which are foreign invaders

Cell-Adhesion Molecules

 Adhere cells to each other and to extracellular material

 cells do not grow or survive normally unless they are mechanically linked to the extracellular material

Glycocalyx

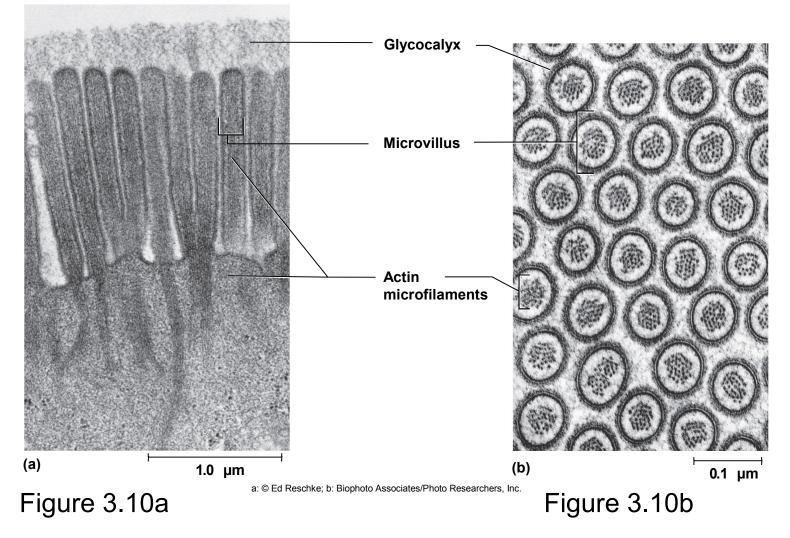
- carbohydrate groups of membrane glycoproteins and glycolipids
 - unique in everyone, but identical twins

Microvilli

- Extensions of membrane (1-2 m)
 - serves to increase cell's surface area
 - best developed in cells specialized in absorption
 - gives 15 40 times more absorptive surface area
- on some cells they are very dense and appear as a fringe "brush border"
 - milking action of actin
 - actin filaments shorten microvilli
 - pushing absorbed contents down into cell

Microvilli

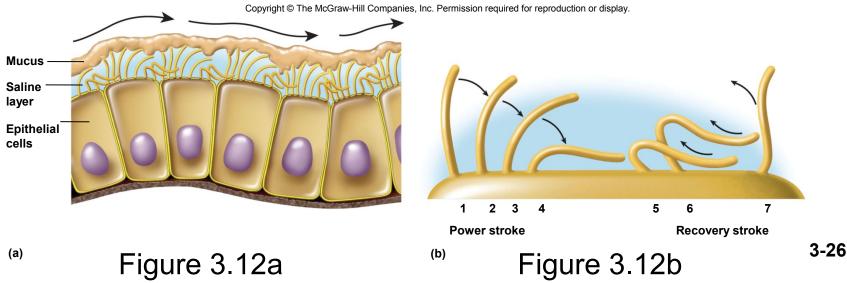
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Actin microfilaments are found in center of each microvilli. 3-25

Cilia

- Hairlike processes 7-10 Im long
 - single, **nonmotile primary cilium** found on nearly every cell
 - "antenna' for monitoring nearby conditions
 - sensory in inner ear, retina, nasal cavity, and kidney
- Motile cilia respiratory tract, uterine tubes, ventricles of the brain, efferent ductules of testes
 - beat in waves
 - sweep substances across surface in same direction
 - power strokes followed by recovery strokes



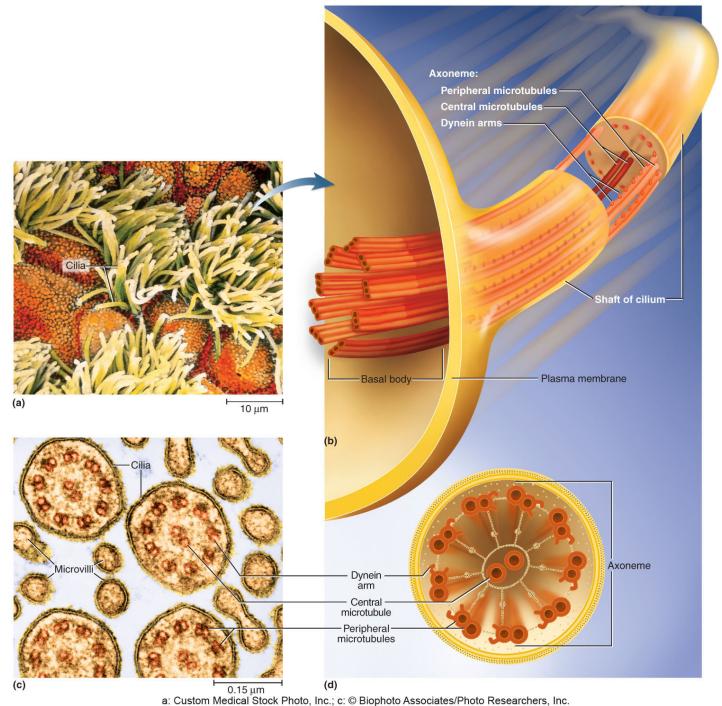
Cross Section of a Cilium

- Axoneme core of cilia that is the structural basis for ciliary movement
- has 9 + 2 structure of microtubules
 - 9 pairs form **basal body** inside the cell membrane
 - anchors cilium
 - dynein arms "crawls" up adjacent microtubule bending the cilia
 - uses energy from ATP

Saline Layer

- chloride pumps pump Cl- into ECF
- Na+ and H2O follows
- cilia beat freely in saline layer

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Cilia & Cystic Fibrosis

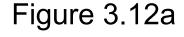
Mucus Saline

layer

Epithelial cells

- Saline layer at cell surface due to chloride pumps move CI- out of cell. Na+ ions and H₂O follow
- Cystic fibrosis hereditary disease in which cells make chloride pumps, but fail to install them in the plasma membrane
 - chloride pumps fail to create adequate saline layer on cell surface
- thick mucus plugs pancreatic ducts and respiratory tract
 - inadequate digestion of nutrients and absorption of oxygen
 - chronic respiratory infections
 - life expectancy of 30

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Flagella

- tail of the sperm only functional flagellum
- whiplike structure with axoneme identical to cilium
 - much longer than cilium
 - stiffened by coarse fibers that supports the tail
- movement is more undulating, snakelike
 - no power stroke or recovery stroke as in cilia

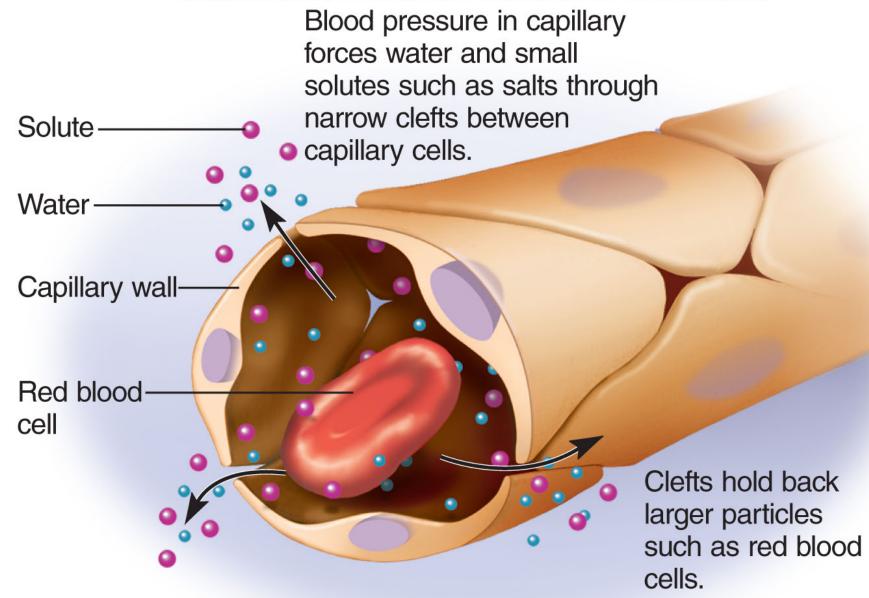
Membrane Transport

- plasma membrane a barrier and a gateway between the cytoplasm and ECF
 - selectively permeable allows some things through, and prevents other things from entering and leaving the cell
- passive transport mechanisms requires no ATP
 - random molecular motion of particles provides the necessary energy
 - filtration, diffusion, osmosis
- active transport mechanisms consumes ATP
 - active transport and vesicular transport
- carrier-mediated mechanisms use a membrane protein to transport substances from one side of the membrane to the other

Filtration

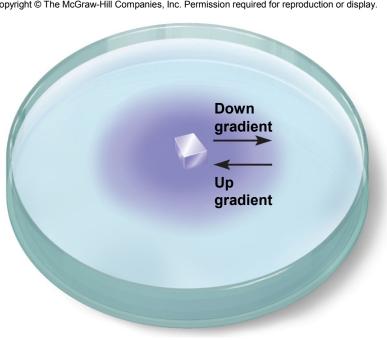
- Filtration process in which particles are driven through a selectively permeable membrane by hydrostatic pressure (force exerted on a membrane by water)
- Examples
 - filtration of nutrients through gaps in blood capillary walls into tissue fluids
 - filtration of wastes from the blood in the kidneys while holding back blood cells and proteins

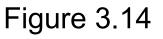
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Simple Diffusion

- Simple Diffusion the net movement of particles from area of high concentration to area of low concentration
 - due to their constant, spontaneous motion
- Also known as movement down the concentration gradient – concentration of a substance differs from one point to another





Diffusion Rates

- Factors affecting diffusion rate through a membrane
 - temperature *temp., *motion of particles
 - molecular weight larger molecules move slower
 - steepness of concentrated gradient *difference, *rate
 - membrane surface area ★area, ★rate
 - membrane permeability *permeability, *rate

Membrane Permeability

Diffusion through lipid bilayer

 Nonpolar, hydrophobic, lipid-soluble substances diffuse through lipid layer

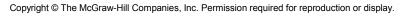
Diffusion through channel proteins

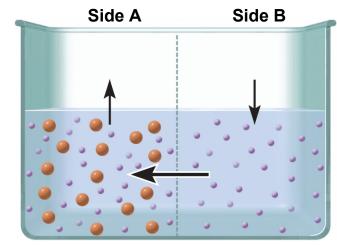
 water and charged, hydrophilic solutes diffuse through channel proteins in membrane

• Cells control permeability by regulating number of channel proteins or by opening and closing gates

Osmosis

- Osmosis flow of water from one side of a selectively permeable membrane to the other
 - from side with higher water concentration to the side with lower water concentration
 - reversible attraction of water to solute particles forms hydration spheres
 - makes those water molecules less available to diffuse back to the side from which they came
- Aquaporins channel proteins specialized for passage of water



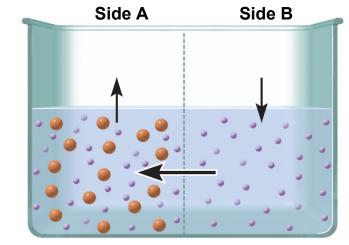


(a) Start

Figure 3.15a

Aquaporins

- Aquaporins channel proteins in plasma membrane specialized for passage of water
 - cells can increase the rate of osmosis by installing more aquaporins
 - decrease rate by removing them
- Significant amounts of water diffuse even through the hydrophobic, phospholipid regions of the plasma membrane



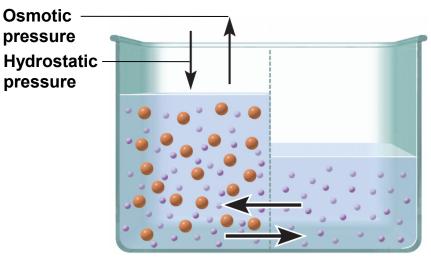
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(a) Start

Figure 3.15a

Osmotic Pressure

- Osmotic Pressure amount of hydrostatic pressure required to stop osmosis
- Osmosis slows due to hydrostatic pressure
- Heart drives water out of capillaries by reverse osmosis – capillary filtration
- Osmolarity measure of osmotic potential



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(b) 30 minutes later

Figure 3.15b

Tonicity

- **Tonicity** ability of a solution to affect fluid volume and pressure in a cell
 - depends on concentration and permeability of solute

Hypotonic solution

- has a lower concentration of nonpermeating solutes than intracellular fluid (ICF)
 - high water concentration
- cells absorb water, swell and may burst (lyse)

Hypertonic solution

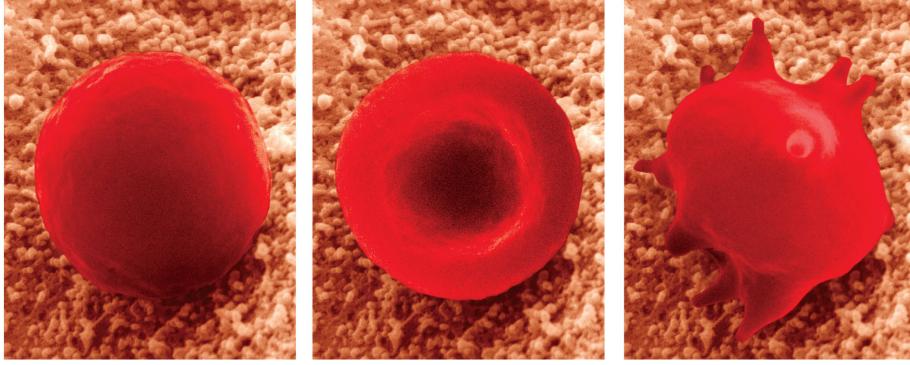
- has a higher concentration of nonpermeating solutes
 - low water concentration
- cells lose water + shrivel (crenate)

Isotonic solution

- concentrations in cell and ICF are the same
- cause no changes in cell volume or cell shape
- normal saline

Effects of Tonicity on RBCs

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(a) Hypotonic

(b) Isotonic

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Figure 3.16a Figure 3.16b Figure 3.16c Hypotonic, isotonic and hypertonic solutions affect the fluid volume of a red blood cell. Notice the crenated and swollen cells.

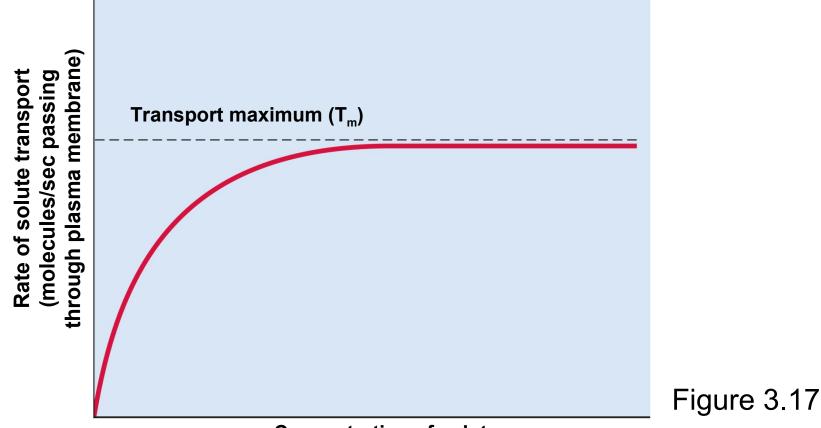
(c) Hypertonic

Carrier-Mediated Transport

- **Transport proteins** in the plasma membrane that carry solutes from one side of the membrane to the other
- Specificity:
 - transport proteins specific for a certain **ligand**
 - solute binds to a specific receptor site on carrier protein
 - differs from membrane enzymes because carriers do not chemically change their ligand
 - simply picks them up on one side of the membrane, and release them, unchanged, on the other
- Saturation:
 - as the solute concentration rises, the rate of transport rises, but only to a point – Transport Maximum (Tm)
- 2 types of carrier mediated transport
 - facilitated diffusion and active transport

Membrane Carrier Saturation

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Concentration of solute

 Transport maximum - transport rate when all carriers are occupied

Membrane Carriers

Uniport

- carries only one solute at a time

Symport

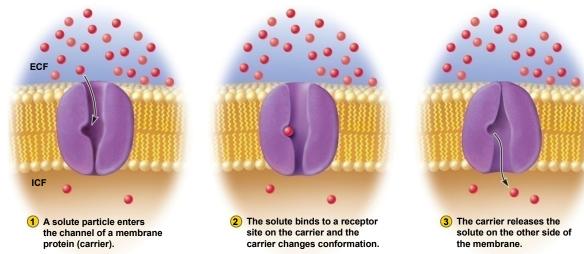
 carries 2 or more solutes simultaneously in same direction (cotransport)

Antiport

- carries 2 or more solutes in opposite directions (countertransport)
- sodium-potassium pump brings in K⁺ and removes Na⁺ from cell
- carriers employ two methods of transport
 - facilitated diffusion
 - active transport

Facilitated Diffusion

- facilitated diffusion carrier-mediated transport of solute through a membrane down its concentration gradient
- does not consume ATP
- solute attaches to binding site on carrier, carrier changes confirmation, then releases solute on other side of membrane



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Figure 3.18

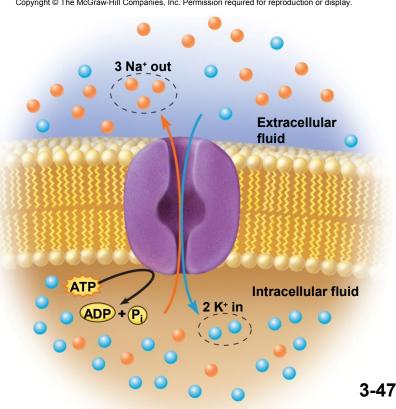
Active Transport

- active transport carrier-mediated transport of solute through a membrane up (against) its concentration gradient
- ATP energy consumed to change carrier
- Examples of uses:
 - sodium-potassium pump keeps K+ concentration higher inside the cell
 - bring amino acids into cell
 - pump Ca²⁺ out of cell

Sodium-Potassium Pump

- each pump cycle consumes one ATP and exchanges three Na+ for two K+
- keeps the K+ concentration higher and the Na+ concentration lower with in the cell than in ECF
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 </sup>
- necessary because Na⁺ and K⁺ constantly leak through membran
 - half of daily calories utilized for Na+ - K+ pump

Figure 3.19



Functions of Na⁺ -K⁺ Pump

Regulation of cell volume

- "fixed anions" attract cations causing osmosis
- cell swelling stimulates the Na⁺- K⁺ pump to

 ❀ion concentration, ❀osmolarity and cell swelling

Secondary active transport

- steep concentration gradient maintained between one side of the membrane and the other – (water behind a dam)
- Sodium-glucose transport protein (SGLT) simultaneously binds Na+ and glucose and carries both into the cell
- does not consume ATP

Heat production

- thyroid hormone increase # of Na+ K+ pumps
- consume ATP and produce heat as a by-product

• Maintenance of a membrane potential in all cells

- pump keeps inside more negative, outside more positive
- necessary for nerve and muscle function

Vesicular Transport

 Vesicular Transport – processes that move large particles, fluid droplets, or numerous molecules at once through the membrane in vesicles – bubblelike enclosures of membrane

- motor proteins consumes ATP

- Endocytosis -vesicular processes that bring material into the cell
 - phagocytosis "cell eating" engulfing large particles
 - pseudopods phagosomes macrophages
 - pinocytosis "cell drinking" taking in droplets of ECF containing molecules useful in the cell
 - pinocytic vesicle
 - receptor-mediated endocytosis particles bind to specific receptors on plasma membrane
 - clathrin-coated vesicle
- **Exocytosis** discharging material from the cell
- Utilizes motor proteins energized by ATP

Phagocytosis or "Cell-Eating"

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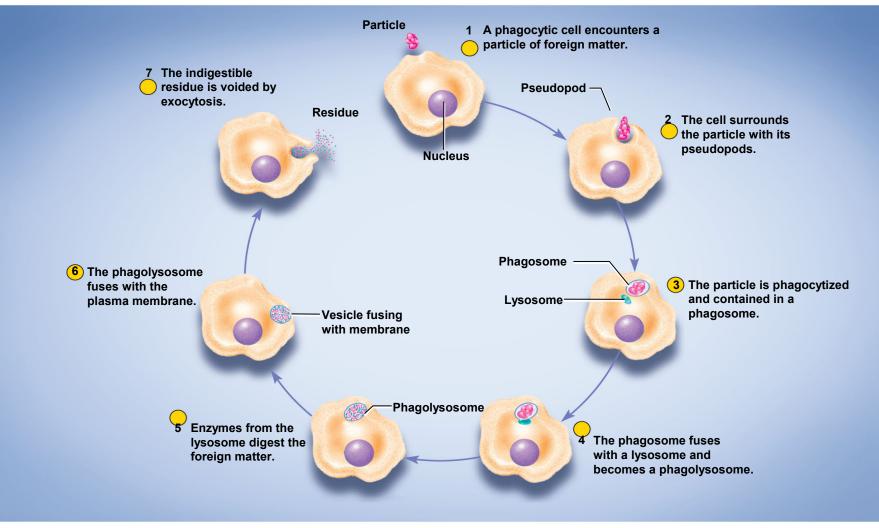


Figure 3.21

Keeps tissues free of debris and infectious microorganisms.

3-50

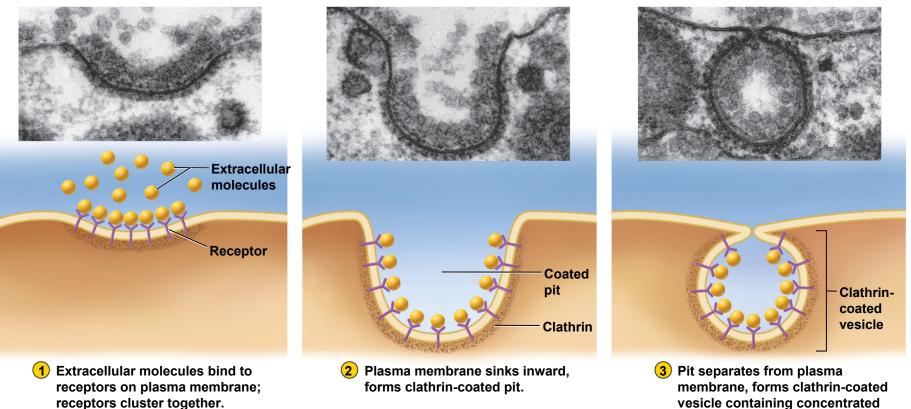
Pinocytosis or "Cell-Drinking"

- Taking in droplets of ECF
 - occurs in all human cells

 Membrane caves in, then pinches off into the cytoplasm as pinocytotic vesicle

Receptor Mediated Endocytosis

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(all): Company of Biologists, Ltd.

Figure 3.22 (1,2 and 3)

3-52

molecules from ECF.

Receptor Mediated Endocytosis

- more selective endocytosis
- enables cells to take in specific molecules that bind to extracellular receptors
- Clathrin-coated vesicle in cytoplasm
 uptake of LDL from bloodstream

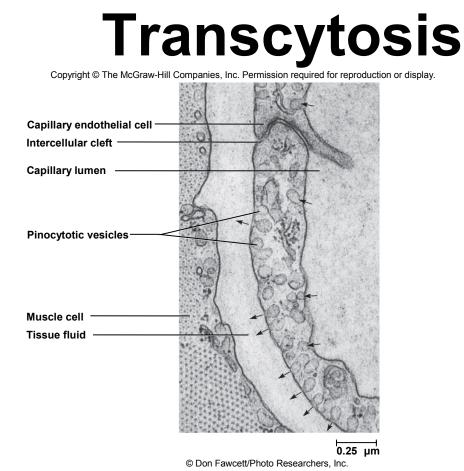


Figure 3.23

- Transport of material across the cell by capturing it on one side and releasing it on the other
- Receptor-mediated endocytosis moves it into cell and exocytosis moves it out the other side

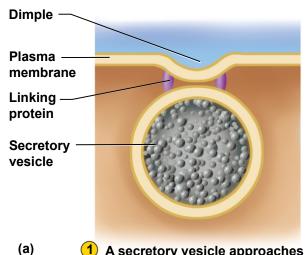
 insulin

Exocytosis

- Secreting material
- replacement of plasma membrane removed by endocytosis

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Fusion pore



 A secretory vesicle approaches the plasma membrane and docks on it by means of linking proteins. The plasma membrane caves in at that point to meet the vesicle.

2 The plasma membrane and vesicle unite to form a fusion pore through which the vesicle contents are released.

Courtesy of Dr. Birgit Satir, Albert Einstein College of Medicine

Secretion

(b)

The Cell Interior

- structures in the cytoplasm
 - organelles, cytoskeleton, and inclusions
 - all embedded in a clear gelatinous **cytosol**
- Organelles internal structures of a cell that carry out specialized metabolic tasks
 - membranous organelles those surrounded by one or two layers of unit membrane
 - nucleus, mitochondria, lysosome, peroxisome, endoplasmic reticulum, and Golgi complex
 - organelles not surrounded by membranes
 - ribosome, centrosome, centriole, basal bodies

Cytoskeleton

- collection of protein filaments
- microfilaments, intermediate filaments, and microtubules

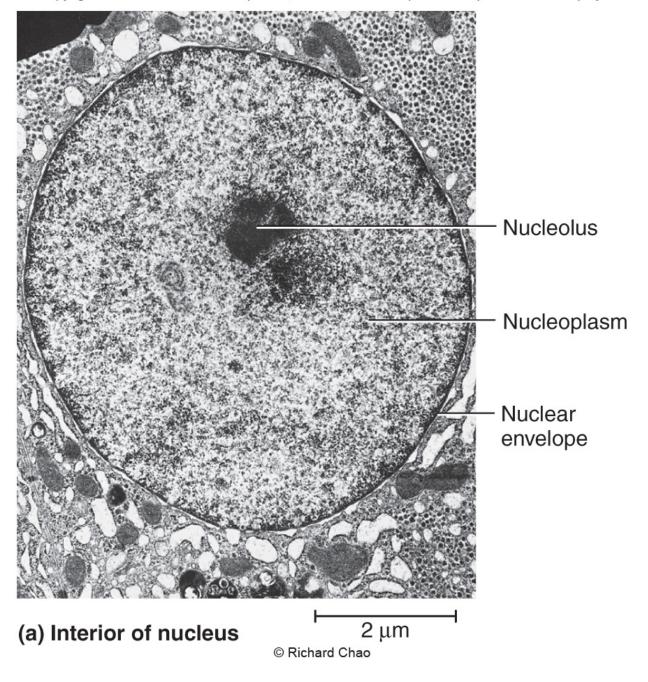
Inclusions

stored cellular components and fat droplets

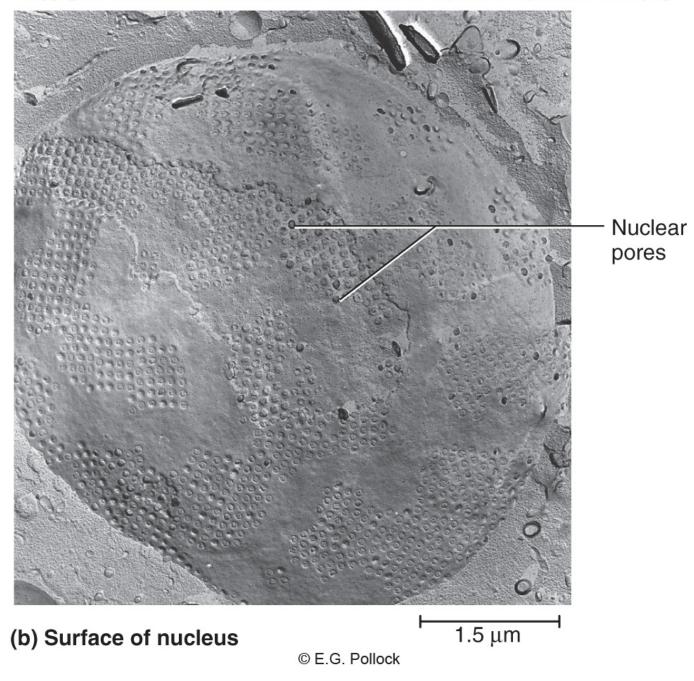
Nucleus

- Largest organelle (5 Im in diameter)
 - most cells have one nucleus
 - a few cells are **anuclear** or **multinucleate**
- **nuclear envelope -** two unit membranes surround nucleus
 - perforated by nuclear pores formed by rings of protein
 - regulate molecular traffic through envelope
 - hold two unit membranes together
 - supported by nuclear lamina
 - web of protein filaments
 - supports nuclear envelope and pores
 - provides points of attachment and organization for chromatin
 - plays role in regulation of the cell life cycle
- **nucleoplasm** material in nucleus
 - chromatin (thread-like matter) composed of DNA and protein
 - nucleoli one or more dark masses where ribosomes are produced

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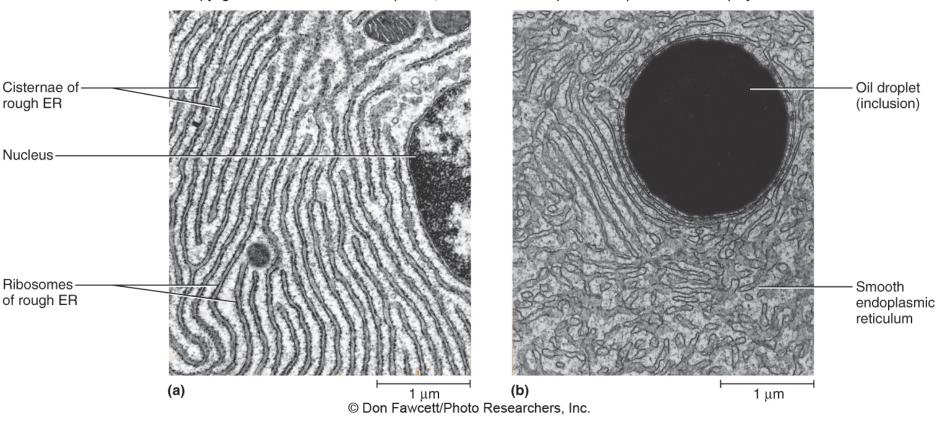


Endoplasmic Reticulum

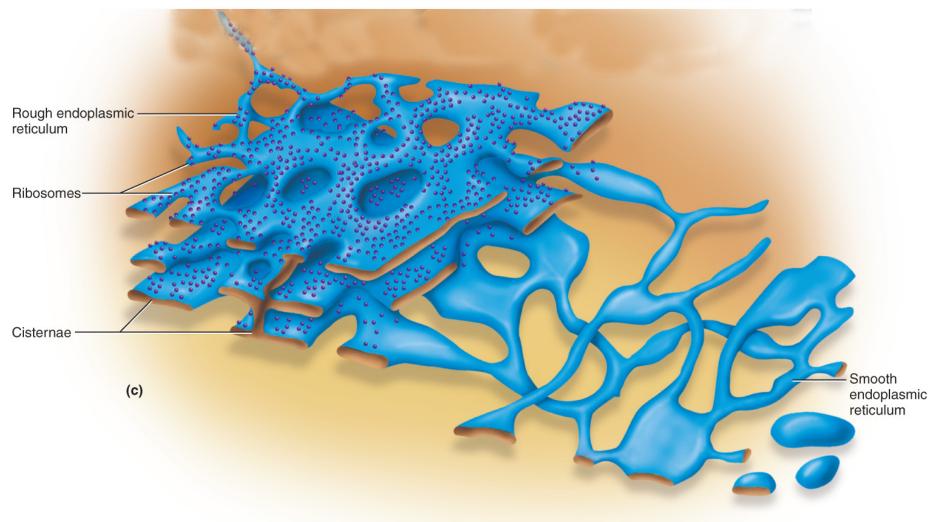
- endoplasmic reticulum system of interconnected channels called cisternae enclosed by unit membrane
- rough endoplasmic reticulum composed of parallel, flattened sacs covered with ribosomes
 - continuous with outer membrane of nuclear envelope
 - adjacent cisternae are often connected by perpendicular bridges
 - produces the phospholipids and proteins of the plasma membrane
 - synthesizes proteins that are packaged in other organelles or secreted from cell

Endoplasmic Reticulum

- smooth endoplasmic reticulum
 - lack ribosomes
 - cisternae more tubular and branching
 - cisternae are thought to be continuous with those of rough ER
 - synthesizes steroids and other lipids
 - detoxifies alcohol and other drugs
 - manufactures all membranes of the cell
- rough and smooth ER are functionally different parts of the same network



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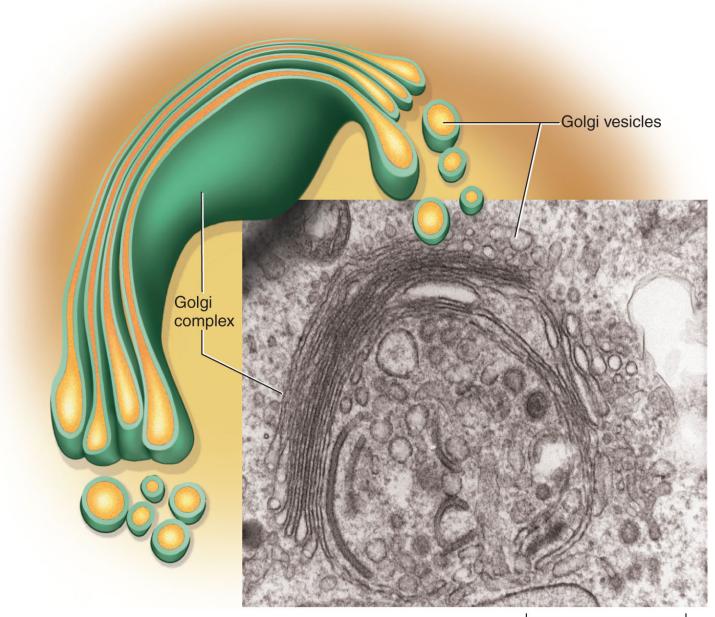
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Ribosomes

- Ribosomes small granules of protein and RNA
 - found in nucleoli, in cytosol, and on outer surfaces of rough ER, and nuclear envelope
- they 'read' coded genetic messages (messenger RNA) and assemble amino acids into proteins specified by the code

Golgi Complex

- **Golgi complex** a small system of cisternae that synthesize carbohydrates and put the finishing touches on protein and glycoprotein synthesis
 - receives newly synthesized proteins from rough ER
 - sorts them, cuts and splices some of them, adds carbohydrate moieties to some, and packages the protein into membrane-bound Golgi vesicles
 - some become lysosomes
 - some migrate to plasma membrane and fuse to it
 - some become secretory vesicles for later release



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Lysosomes

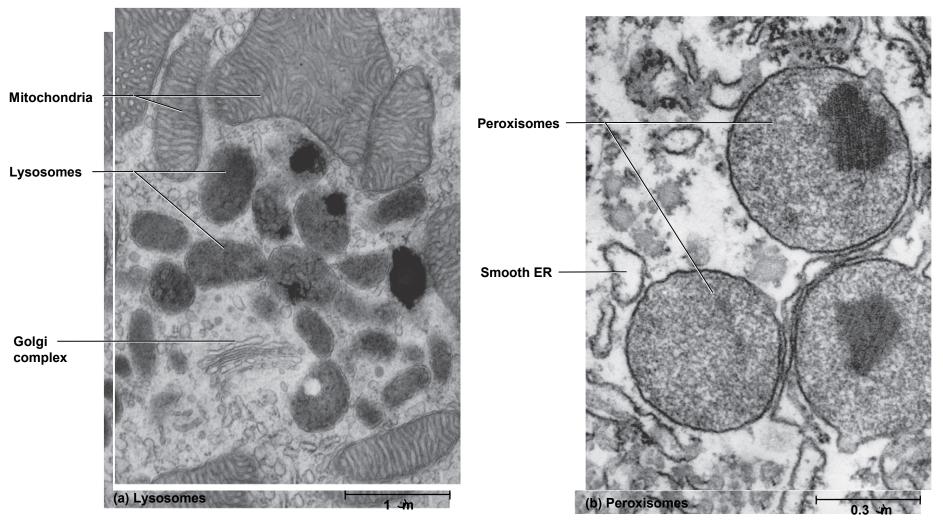
- Lysosomes package of enzymes bound by a single unit membrane
 - extremely variable in shape

Functions

- intracellular hydrolytic digestion of proteins, nucleic acids, complex carbohydrates, phospholipids, and other substances
- autophagy digest and dispose of worn out mitochondria and other organelles
- autolysis 'cell suicide' some cells are meant to do a certain job and then destroy themselves

Lysosomes and Peroxisomes

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Figure 3.28a

Figure 3.28b

Peroxisomes

- Peroxisomes resemble lysosomes but contain different enzymes and are not produced by the Golgi complex
- General function is to use molecular oxygen to oxidize organic molecules
 - these reactions produce hydrogen peroxide (H2O2)
 - catalase breaks down excess peroxide to H2O and O2
 - neutralize free radicals, detoxify alcohol, other drugs, and a variety of blood-borne toxins
 - breakdown fatty acids into acetyl groups for mitochondrial use in ATP synthesis
- In all cells, but abundant in liver and kidney

Mitochondrion

- mitochondria organelles specialized for synthesizing ATP
- variety of shapes spheroid, rodshaped, kidney bean-shaped, or threadlike
- surrounded by a double unit membrane
 - inner membrane has folds called cristae
 - spaces between cristae are called matrix
 - matrix contains ribosomes, enzymes used for ATP synthesis, small circular DNA molecule – mitochondrial DNA (mtDNA)
- "Powerhouses" of the cell
 - energy is extracted from organic molecules and transferred to ATP

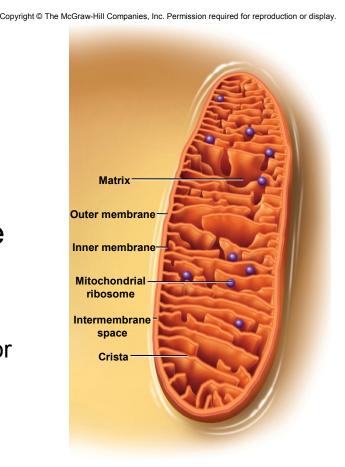


Figure 3.29b

Mitochondrion

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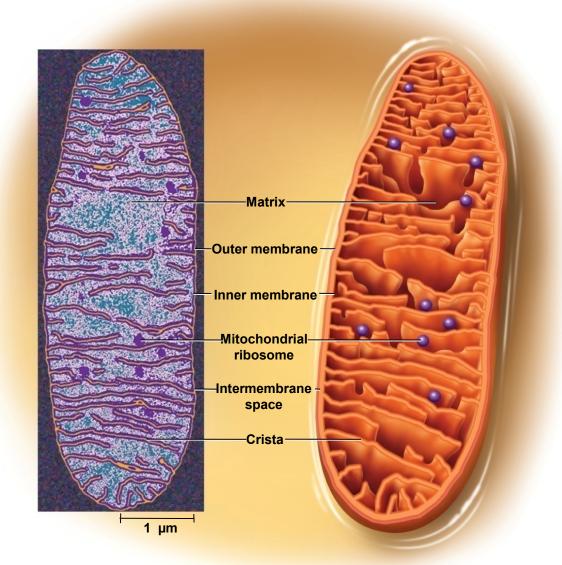


Figure 3.29a,b

Evolution of Mitochondrion

- It is a virtual certainty that mitochondria evolved from bacteria that invaded another primitive cell, survived in the cytoplasm, and became permanent residents
 - its two unit membranes suggests that the original bacterium provided the inner membrane, and the host cell's phagosome provided the outer membrane
 - mitochondrial ribosomes more like bacterial ribosomes
 - has its own mtDNA
 - small circular molecule resembling bacterial DNA
 - replicates independently of nuclear DNA
 - when a sperm fertilizes the egg, any mitochondria introduced by the sperm are usually destroyed, and only those provided by the egg are passed on to the developing embryo
 - mitochondrial DNA is almost exclusively inherited through the mother
 - mutates more readily than nuclear DNA
 - no mechanism for DNA repair
 - produces rare hereditary diseases
 - mitochondrial myopathy, mitochondrial encephalomyopathy, and others

Centrioles

- centriole a short cylindrical assembly of microtubules arranged in nine groups of three microtubules each
- two centrioles lie perpendicular to each other within a small clear area of cytoplasm - centrosome

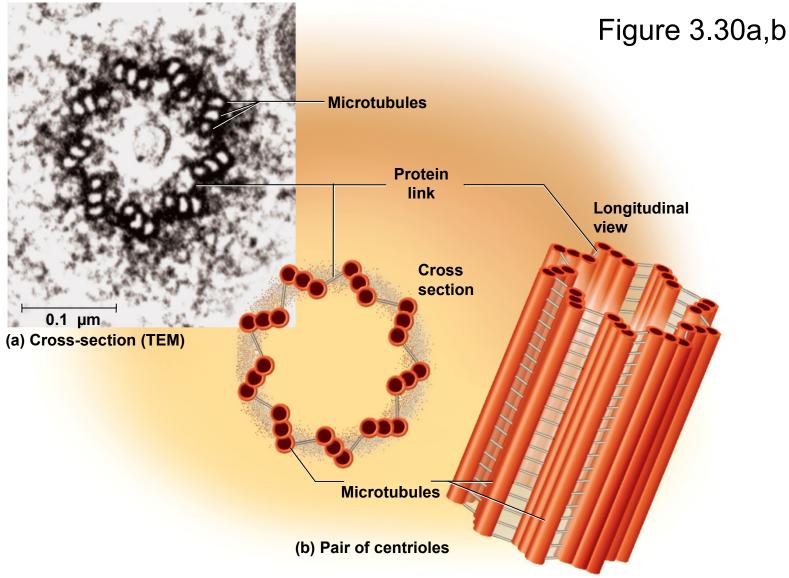
 play role in cell division

cilia and flagella formation

- each basal body of a cilium or flagellum is a single centriole oriented perpendicular to plasma membrane
- basal bodies originate in *centriolar organizing center*
 - migrates to plasma membrane
- two microtubules of each triplet elongate to form the nine pairs of peripheral microtubules of the **axoneme**
- cilium reaches full length in less than one hour

Centrioles

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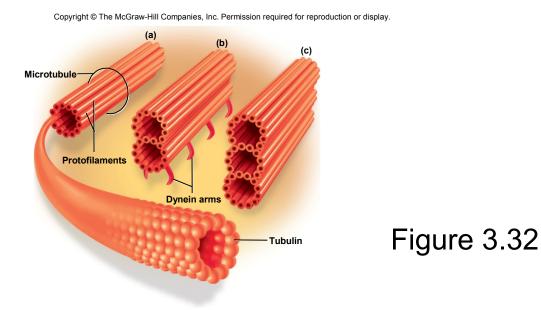


Cytoskeleton

- Cytoskeleton collection of filaments and cylinders
 - determines shape of cell, lends structural support, organizes its contents, directs movement of substances through the cell, and contributes to the movements of cell as a whole.
- Composed of:
 - microfilaments made of protein actin
 - form network on cytoplasmic side of plasma membrane called the terminal web
 - provides physical support for phospholipid bilayer
 - actin supports microvilli and produces cell movements
 - intermediate fibers thicker and stiffer than microfilaments
 - resist stresses placed on cell
 - · participate in junctions that attach some cells to their neighbors
 - line nuclear envelope and form cagelike nuclear lamina that encloses DNA

Microtubules

- microtubule made of the protein tubulin
- microtubules radiate from centrosome and hold organelles in place, form bundles that maintain cell shape and rigidity, and act somewhat like railroad tracks
 - motor proteins 'walk' along these tracks carrying organelles and other macromolecules to specific locations in the cell
- not permanent structures, they come and go moment by moment





Cytoskeleton

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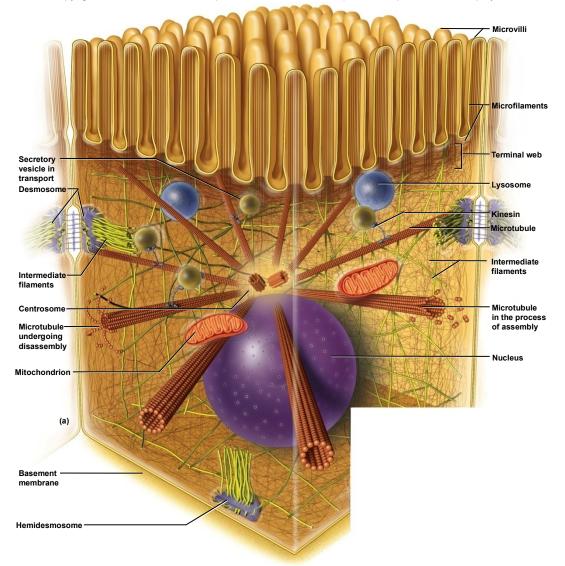
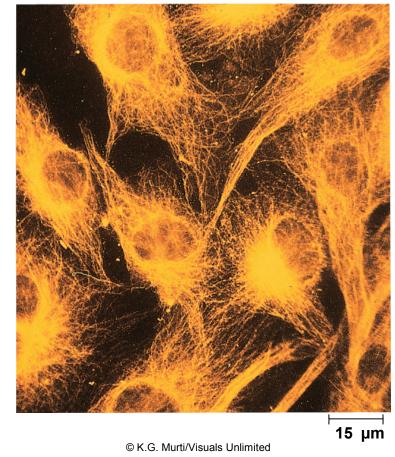


Figure 3.31a

EM and Fluorescent Antibodies demonstrate Cytoskeleton

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(b)

Figure 3.31b