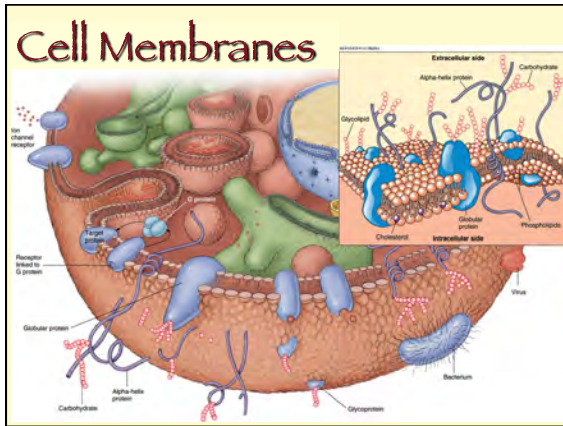


Cell Membranes



What does a cell need?

- **Selective isolation from environment (plasma membrane)**
- **Energy (ATP)** – [to be discussed in future lecture]
- Instructions (DNA)
- Machinery to carry out instructions and regulate processes (proteins)
- **Compartmentalization of incompatible or specialized activities (organelles)**

Cell Membrane Function

- Boundary between internal and external environments
- Selectively permeable - controls what goes in and out of cell or organelle
- Attachment to extracellular surfaces or to other cells (and organelles to cytoskeleton)
- Self or species recognition
- Cell to cell communication
- Lipid metabolism
- Localization of fixed or sequential processes

Membrane Phospholipid Bilayer

- Phospho-lipid bilayer forms the essential backbone of cellular membranes.

The diagram shows a cross-section of a phospholipid bilayer. The hydrophilic heads are oriented towards the water on both sides, while the hydrophobic tails are oriented towards each other, away from the water. A symbol for a phospholipid is shown with its head and tails.

Synthesis of membrane lipids

- Phospholipid synthesis in smooth endoplasmic reticulum

Copyright © The McGraw-Hill Companies, Inc. Permission required for reproduction or display.

- 1 In the cytosol, fatty acids are activated by the attachment of a CoA molecule.
- 2 The activated fatty acids bind to glycerol phosphate and are inserted into the cytosolic leaflet of the ER membrane via acyl transferase.
- 3 The phosphate is removed by a phosphatase enzyme.
- 4 A choline linked to phosphate is attached via choline phosphotransferase.
- 5 Flippase transfer some of the phospholipids to the other leaflet.

The diagram shows the chemical structures of the reactants: 2 fatty acids, 2 activated molecules (fatty acid-CoA), and glycerol-phosphate. The reaction involves the enzyme acyl transferase in the cytosol, followed by phosphatase and choline phosphotransferase in the ER lumen. The final product is a phospholipid with a choline head group. A flippase is also shown moving lipids between the leaflets.

Synthesis of membrane lipids

- Distribution of membrane lipids from sER to other organelles – Three mechanisms

Copyright © The McGraw-Hill Companies, Inc. Permission required for reproduction or display.

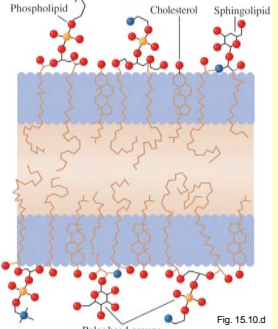
The diagram illustrates three mechanisms for lipid transport: (a) Lateral diffusion, where lipids move from the ER membrane to other membranes; (b) Vesicle transport, where lipids are packaged into vesicles that fuse with other organelles; and (c) Lipid exchange proteins, which retrieve lipids from other organelles and insert them into the ER membrane. The diagram also shows the ER membrane and its connection to the nucleus and mitochondria.

Cell Membranes

Synthesis of membrane lipids

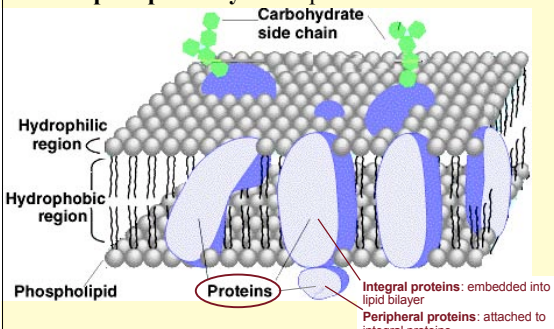
Types of membrane lipids:

- **Phospholipids**
 - Major component of lipid bilayer
- **Cholesterol**
 - Contributes to rigidity / flexibility of membrane
- **Sphingolipids**
 - Two fatty acids condensed onto a serine, instead of a glycerol
 - Usually with longer fatty chains than on phospholipids
 - Polar head may be modified with a phosphate or carbohydrate (glycolipid)
 - Rafts of sphingolipids migrate along the membrane



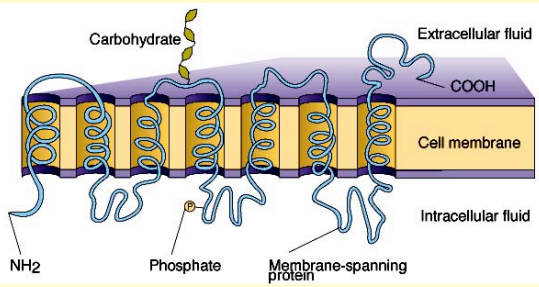
Plasma Membrane Proteins

- **Phospholipid bilayer** has proteins embedded in it.

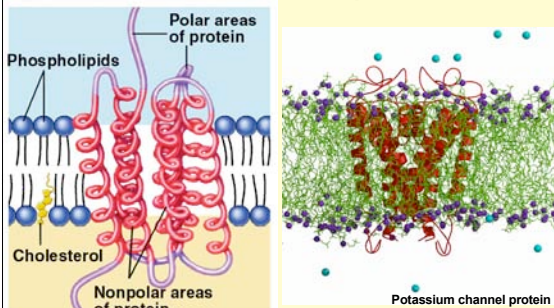


Proteins can insert into lipid bilayer

- Protein domains can be hydrophobic or hydrophilic



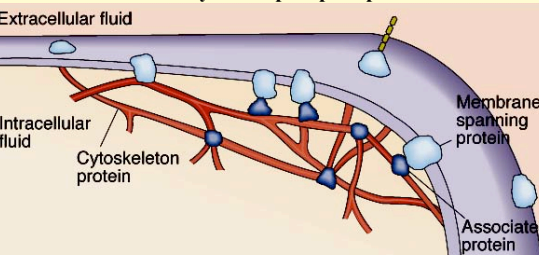
Transmembrane proteins



Multiple hydrophobic domains contribute to intramembrane function

Plasma Membrane Proteins

Provide functionality to the phospholipid barrier



Functions of Plasma Membrane Proteins

Outside			
Plasma membrane			
Inside			
	Transporter	Enzyme	Cell surface receptor
	Cell surface identity marker	Cell adhesion	Attachment to the cytoskeleton

- Gates
- Pumps
- Receptors
- Ligands
- Anchors
- Junctions
- Fixed enzymes
- Lipophilic enzymes

Cell Membranes

Fluid Mosaic Model of Membrane Structure

- **Mosaic:** patchy, non-uniform distribution of proteins
 - Different regions or sides of the same cell may have different functions
- **Fluid:** distribution is dynamic and changeable

Fluid Mosaic Model of Membrane Structure

proteins can move laterally within membrane

Importing proteins to organelles

Membrane-enclosed organelles import proteins by one of three mechanisms. All of these processes require energy. The protein remains folded during the transport steps in mechanisms 1 and 3, but usually has to be unfolded in mechanism 2.

Figure 15-5 Essential Cell Biology, 2/e. © 2004 Garland Science

Membrane Carbohydrates

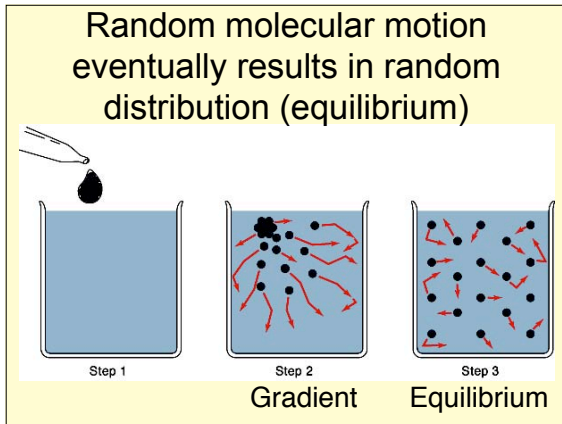
- As glycoprotein or glycolipid
- Added onto extracellular surface groups

Movement of Molecules Across Membranes

Passive Transport (Diffusion)

- Net movement of molecules from a region of high concentration to a region of low concentration
- ✓ Caused by random (Brownian) movements of molecules
- ✓ (Increase entropy)
- ✓ Each type of molecule follows its own concentration gradient
- ✓ At equilibrium, movement is equal in both directions

Cell Membranes



Factors that affect Rate of Diffusion

- Concentration gradient
 - Difference in concentration between two points
- Temperature (molecular movement)
- Permeability of the membrane / medium
- Available surface area of membrane
- Distance across which diffusion must occur
- Solvent state (gas > liquid > semisolid)

FICK'S LAW *Adolf Fick, 1858*

Fick's law of diffusion of a gas across a fluid membrane:

Rate of diffusion = $KA(P_2 - P_1) / D$

Wherein:

- K = a *temperature-dependent* diffusion constant.
- A = the surface area available for diffusion.
- $(P_2 - P_1)$ = The difference in concentration (partial pressure) of the gas across the membrane.
- D = the distance over which diffusion must take place.

Increased exchange rate by increased surface area

- **Microvilli**

Figure 6.26 0.25 μm

Concentration

= Number of solutes in a given volume

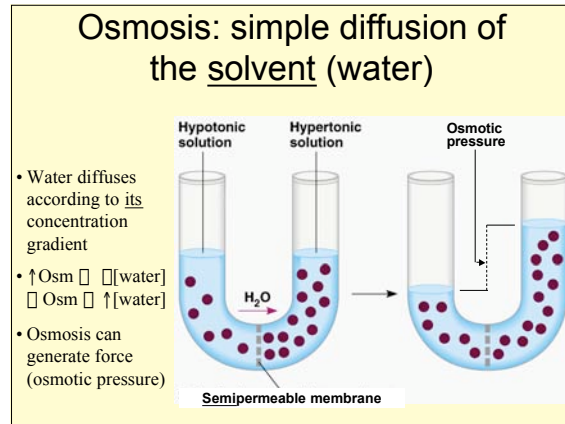
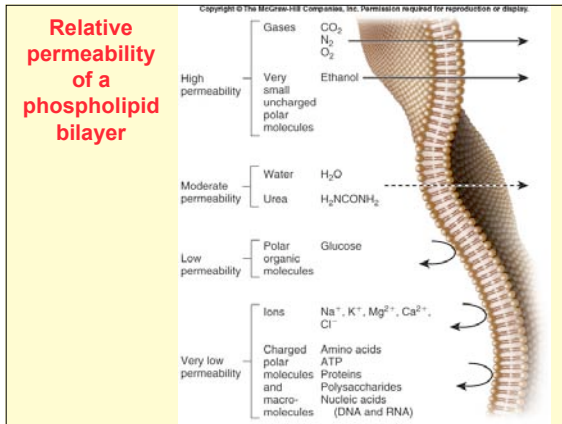
- Examples
 - Moles per liter (molar = *M*)
 - Grams per 100ml (g%)
 - Nanograms per milliliter (ng/ml)
 - Parts per thousand (ppt)
- Osmolarity:
 - the sum of **all** solutes in a given volume
 - in moles per liter (*Osm*)

Simple (non-selective) diffusion across cell membranes

Nonpolar solutes dissolve through membrane

- O₂
- CO₂
- Fat soluble hormones (steroids)
- Urea
- Fat soluble vitamins
- Other small, fat soluble molecules

Cell Membranes

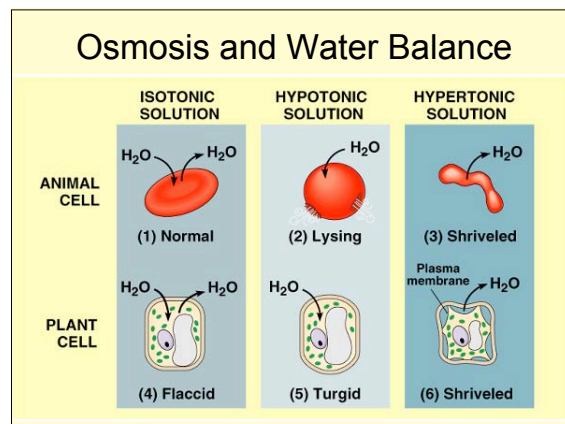
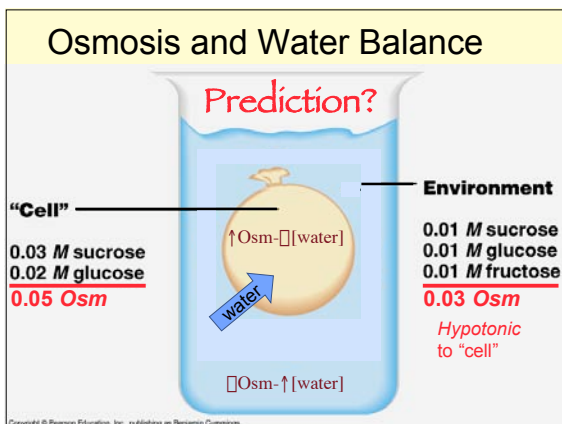


Osmolarity & Osmotic Pressure

- Osmolarity (Osm):** the sum of all solutes in a given volume (moles per liter)
 - 1 M glucose solution = 1 Osm
 - 1 M glucose / 1 M fructose / 1 M ribose solution = 3 Osm
 - 1 M NaCl solution = 1 M Na⁺ / 1 M Cl⁻ = 2 Osm
- Isosmotic:** two solutions with the same Osm
- Hyposmotic:** a solution with a lower Osm than another
- Hyperosmotic:** a solution with a higher Osm than another
- Remember:**
 - ↑ Osm → ↓ [water]
 - ↓ Osm → ↑ [water]

Osmolarity & Osmotic Pressure

- Osmolarity (Osm):** the sum of all solutes in a given volume (moles per liter)
- Osmotic Pressure (P_{Osm}):**
 - Force generated by osmosis
 - Measure of the tendency to take on water by osmosis
- Isotonic:** two solutions with the same P_{Osm}
- Hypotonic:** a solution with a lower P_{Osm} than another
 - I.e., loses water to the other solution
- Hypertonic:** a solution with a higher P_{Osm} than another
 - I.e., takes water from the other solution
- For an isosmotic solution to be isotonic, the membrane must be equally permeable (or equally impermeable) to all solutes*
 - All isotonic solutions are isosmotic.
 - But not all isosmotic solutions are isotonic.



Osmotic Swelling

- Mechanisms to resist excessive swelling in hypotonic environments

(A) ANIMAL CELL (B) PLANT CELL (C) PROTOZOAN

Figure 10-17 Essential Cell Biology, 2/e, © 2004 Garland Science

Selective permeability

- Except for water and small nonpolar solutes, permeability of cell membranes is **selective and regulated**.
- Permeability determined by **transporter proteins**.
 - Channels and carriers are solute specific
 - If no transporter, than that solute cannot cross membrane
- (Artificial membranes are only semipermeable —i.e., only discriminate based upon molecular size.)

Types of cellular transport

- Passive transport:** driven by Brownian motion
 - Simple diffusion & osmosis
 - Facilitated diffusion (carrier mediated passive transport)
- Active transport:** requires chemical energy (ATP)
 - Carrier mediated
 - Can transport against concentration gradient

Passive transport Active transport

Carrier mediated transport

- Transporters are *binding proteins*
- subject to
 - Specificity
 - Competition
 - Saturation

Rate of transport into cell

Extracellular substrate concentration

Saturation: when all available transporter proteins are in use

Facilitated Diffusion

- Carrier-mediated *passive* transport—“gates” & “channels”

Outside of cell

- Movement from high to low concentration, only when gate is open
- Gate may open/close in response to
 - Chemical signal
 - Cell voltage
 - Mechanical distortion

Facilitated Diffusion

Glucose binding site
Hydrophobic
Hydrophilic
Charged amino acids

Exterior
Cytoplasm

Glucose channel

- May be gated
 - Most tissues
 - Opens in response to insulin
- Or ungated
 - Brain tissue

Cell Membranes

Facilitated Diffusion

• Multidrug Transporter Mechanism

Osmosis may be both simple and facilitated

Aquaporins (water channels) speed water movement.

Active Transport

- Carrier mediated – “pumps”
- Active:** requires **ATP**
- Can force movement against concentration gradient
- Creates concentration gradient
- (creates order/ decreases entropy)

Facilitated diffusion Active transport

Active Transport

- Solute binds to carrier protein
- Binding triggers ATP hydrolysis, transfers phosphate to carrier
- Phosphorylation produces change in shape of carrier
- Change in shape causes carrier to move solute

Difference in concentration is maintained by selective permeability of membrane

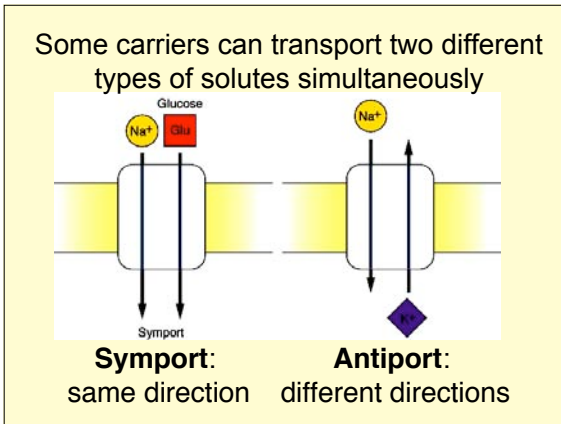
- Cytosol relatively high in K^+ , protein, organic-phosphates
- Low in Na^+ , Ca^{2+} , Cl^-

Figure 11.11

Difference in concentration is maintained by selective permeability of membrane

- Cytosol relatively high in K^+ , protein, organic-phosphates
- Low in Na^+ , Ca^{2+} , Cl^-

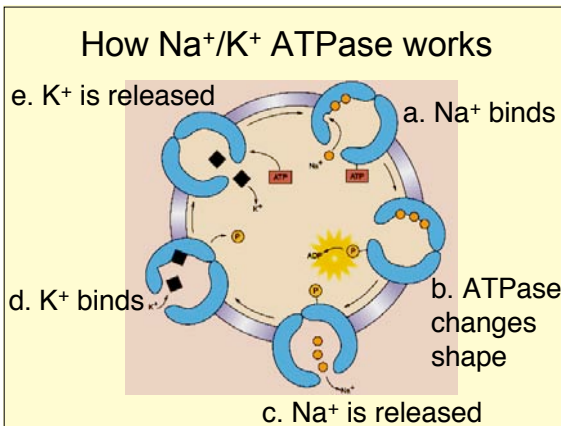
Cell Membranes



Na⁺/K⁺ ATPase:

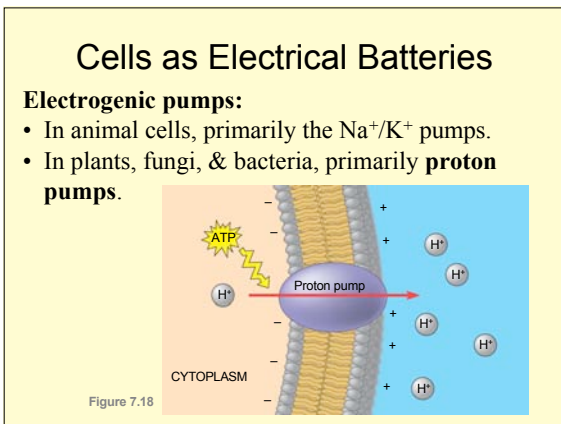
very important active transporter

- Antiport
 - Pumps 3 Na⁺ out
 - Pumps 2 K⁺ in
 } per each ATP used
- Na⁺/K⁺ ATPase used to:
 - Maintain ion gradients
 - Create electrical potential (inside of cell negatively charged relative to outside).



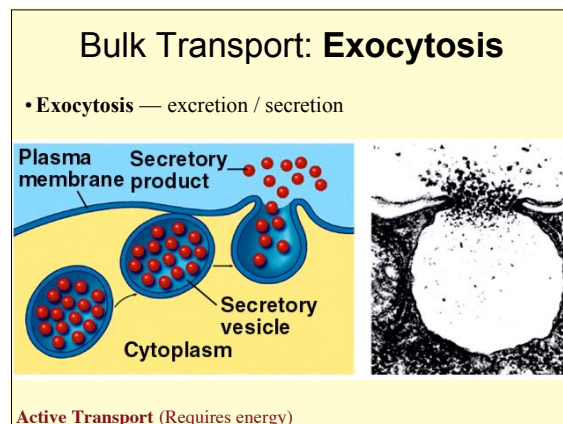
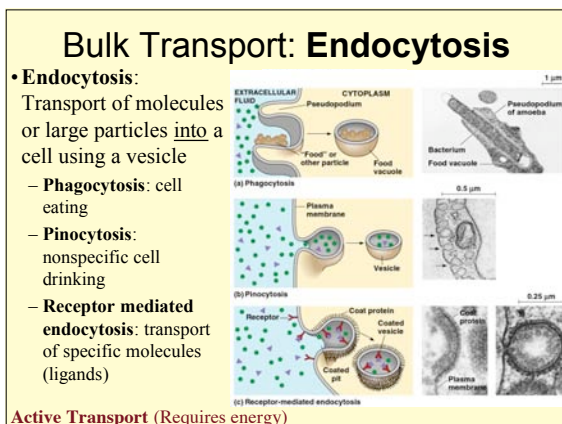
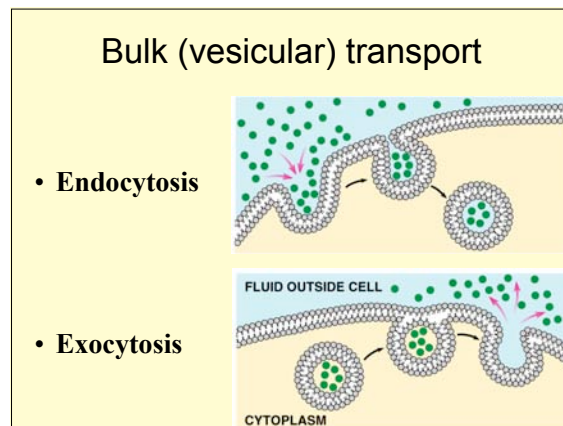
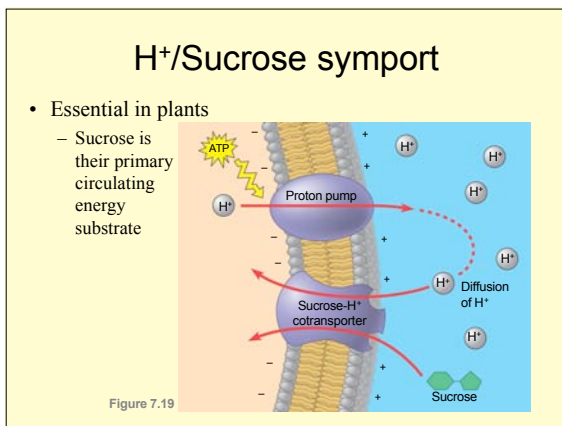
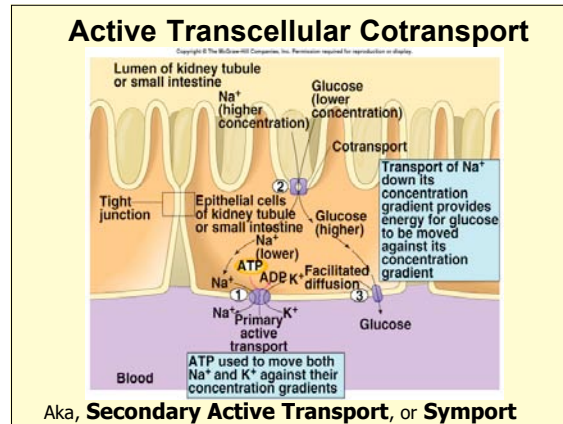
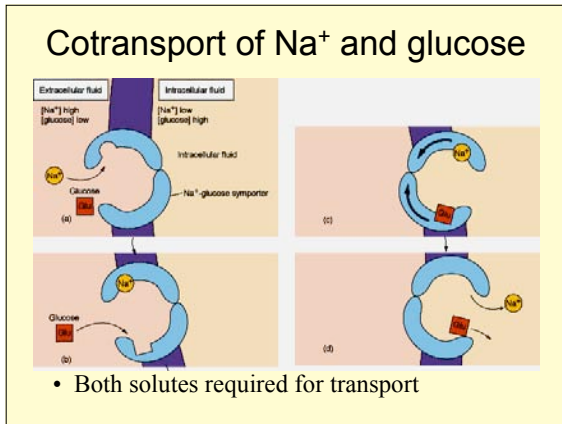
Cells as Electrical Batteries

- **Electrogenic pumps:** Active transport □ chemical gradients of ions □ electrical gradients.
- Electrical gradient produces a membrane potential.
- Inside of the cell is negative relative to the outside of the cell.



Cotransport, or secondary active transport

- Carrier protein does not directly use ATP
- But ATP required to create the gradient by other pumps
- Solute "A" transported by diffusion with the created gradient
- Solute "B" moved against gradient by "piggy-backing" with solute "A"
- Example: Na⁺ and glucose symport
 - Na⁺ diffuses
 - Glucose actively transported



Cell Membranes

Cells Eat and Spit Out: Endo- and Exocytosis

Paramecium

White blood cell

The diagram shows a Paramecium engulfing a microbe through phagocytosis. In a white blood cell, a phagocytic vacuole fuses with lysosomes to kill and digest microbes, followed by exocytosis of debris.

Cell Surface

- Carbohydrates (on glycoproteins and glycolipids) give membranes "sidedness"
 - Membrane-bound carbs for cell recognition
 - Secreted glycoproteins coat outer surface of cell

Labels in diagram: Transmembrane glycoproteins, Secretory protein, Golgi apparatus, Vesicle, Attached carbohydrate, Glycolipid, ER lumen, Plasma membrane: Cytoplasmic face, Extracellular face, Transmembrane glycoprotein, Secreted protein, Membrane glycolipid.

Legend:

- Organelle cytoplasmic (outer) face → plasma membrane cytoplasmic (inner) face
- Organelle lumen (inner) face → plasma membrane extracellular (outer) face

Figure 7.9

Cell Surface Junctions

Labels in diagram: Cell surface coating, Tight junction, Anchoring junction, Communicating junction, Plasma membranes of adjacent cells, Extracellular matrix.

Junctions

- Tight**
 - Restrict fluid passing around cells
- Anchoring**
 - (desmosomes)
 - Join adjacent cytoskeletons
- Communicating (gap)**
 - Allow passage of small molecules
 - Esp. in
 - Plants
 - Embryos
 - Electrochemically coupled cells

Cell Surface Junctions & Walls

Labels in diagram: Plant cell walls, Vacuole, Cytosol, Plasma membrane, Plant cell wall layers: Middle lamella, Primary wall, Secondary wall, Plasmodesma, Primary wall, CELL 1, Three layers of secondary wall, Middle lamella, CELL 2.

- Walls of secreted extracellular polysaccharide (cellulose or chitin)
- Plasmodesmata form gap (communicating) junctions between cells

Extracellular Matrix

Labels in diagram: Proteoglycan molecule, Proteoglycan complex, Polysaccharide molecule, Collagen fiber, Plasma membrane, Integrin, Microfilaments of cytoskeleton, CYTOPLASM.

- No walls (animal cells)
- Matrix or basement membrane of secreted proteins/glycoproteins.

Extracellular Matrix

Major matrix proteins: Collagen and Elastin

Labels in diagram: 50 nm, short section of a collagen fibril, collagen molecule 300 x 1.5 nm, 1.5 nm, elastic fiber, RELAX, STRETCH, single elastin molecule, cross-link.

(A) Collagen is a triple helix formed by three extended protein chains that wrap around one another. Many rodlike collagen molecules are cross-linked together in the extracellular space to form collagen fibrils that have the tensile strength of steel.

(B) Elastin polypeptide chains are cross-linked together to form rubberlike, elastic fibers. Each elastin molecule uncoils into a more extended conformation when the fiber is stretched and will recoil spontaneously as soon as the stretching force is relaxed.

Figure 4-23: Essential Cell Biology, 2/e. © 2004 Garland Science