BCM 226 LECTURE
SALEM CITY, A.J
BIOLOGICAL MEMBRANE

- Biological membranes are composed of proteins associated with a lipid bilayer matrix.
- They are the molecular gateway to the cell.
- Viewed under electron microscope, plasma membrane is a trilaminar structure.
Phospholipid bilayer
Membrane Components

- Phospholipids
- Cholesterol
- Proteins
- Glycocalyx
- Cytoskeleton
- Peripheral protein
- Integral protein

- Phospholipid bilayer
- Cholesterol
- Proteins
- Glycocalyx
MEMBRANE FUNCTION

- Compartmentalization and Protective barrier
- Regulate transport in and out of cell
- Allow cell recognition e.g. cell recognition protein (MHC)
- Provide anchoring sites for filaments of cytoskeleton and scaffold for biochemical reactions
- Provide a binding site for enzymes (receptor)
- Intercellular interaction (interlocking surface or junctions connector)
- Contains the cytoplasm
- Signal transduction
- Enzymatic activity
Cell recognition
Membrane transport may be mediated by the carrier proteins e.g GLUT 2
Binding site for enzyme
Anchoring and scaffold
Intercellular interaction
Signal transduction
For example, the membrane protein, adenylyl cyclase, is involved in ATP metabolism.

Cholera bacteria release a toxin that interferes with the proper functioning of adenylyl cyclase, thus making sodium ion and water leave intestinal cells and the individual dies from severe diarrhea and dehydration.
CELL COMMUNICATION

• Cell communication is very essential for multicellular organisms. For instance, response to pain signals by the muscle cells.
• Why do cells communicate?
• How do cell communicate?
• Cells communicate through any of four basic mechanisms, depending primarily on the distance between the signaling and responding cells.
• In addition, some cells send signals that bind to specific receptors on their own plasma membrane. This is called autocrine signaling which is believed to play an important role in reinforcing developmental process.
Autocrine signaling
• External signals on the cell surface are converted into cellular responses by signal transduction pathways.
• These signals are in form of chemical messengers.
• A hormone is a chemical released by a cell in one part of the body, that sends out messages that affect cells in other parts of the organism.
Types of cell signaling

Direct contact

Paracrine signaling

Synaptic signaling

Endocrine signaling
Paracrine signaling

- Paracrine signals are released by cells into the extracellular fluid in their neighborhood and act locally (short distant). E.g. PGE1

Endocrine signaling: hormone produced in endocrine glands are secreted into the bloodstream and are often distributed widely throughout the body.

Direct contact: Cells that maintain an intimate membrane-to-membrane interface can engage in contact-dependent signaling.

Synaptic signals are transmitted along axons to remote target cells.
Overview of cell signaling

- **Extracellular Fluid**
  - Receptor
  - Signal molecule

- **Plasma membrane**
  - Reception
  - Transduction
  - Relay molecules in a signal transduction pathway

- **Cytoplasm**
  - Response
  - Activation of cellular response
• **Reception** occurs when a signal molecule binds to a receptor protein, causing a conformational change to occur.

• **Transduction:** The binding of the signal molecule alters the receptor protein in some way.

• The signal usually starts a cascade of reactions known as a signal transduction pathway.

• Multistep pathways can amplify a signal.
Response

• Cell signaling leads to regulation of cytoplasmic activities or transcription
  ➢ Signaling pathways regulate a variety of cellular activities
Hormone Receptor

- Nuclear receptor: estrogen
- Cytoplasmic receptors: testosterone and thyroid hormones
- Cell surface receptor: peptide hormone and catecholamines
NUCLEAR RECEPTOR

1. Hormone (H), carried to the target tissue on serum binding proteins, diffuses across the plasma membrane and binds to its specific receptor protein (Rec) in the nucleus.

2. Hormone binding changes the conformation of Rec; it forms homo- or heterodimers with other hormone-receptor complexes and binds to specific regulatory regions called hormone response elements (HREs) in the DNA adjacent to specific genes.

3. Binding regulates transcription of the adjacent gene(s), increasing or decreasing the rate of mRNA formation.

4. Altered levels of the hormone-regulated gene product produce the cellular response to the hormone.
Steroid hormones bind to intracellular receptors

1. The steroid hormone testosterone passes through the plasma membrane.
2. Testosterone binds to a receptor protein in the cytoplasm, activating it.
3. The hormone-receptor complex enters the nucleus and binds to specific genes.
4. The bound protein stimulates the transcription of the gene into mRNA.
5. The mRNA is translated into a specific protein.
**CELL SURFACE RECEPTOR:**

**Receptor Tyrosine Kinase**

- **Insulin receptor binds insulin and undergoes autophosphorylation on its carboxyl-terminal Tyr residues.**

- **Insulin receptor phosphorylates IRS-1 on its Tyr residues.**

- **SH2 domain of Grb2 binds to \(\text{P-Tyr}\) of IRS-1. Sos binds to Grb2, then to Ras, causing GDP release and GTP binding to Ras.**

- **Activated Ras binds and activates Raf-1.**

- **Raf-1 phosphorylates MEK on two Ser residues, activating it. MEK phosphorylates ERK on a Thr and a Tyr residue, activating it.**

- **ERK moves into the nucleus and phosphorylates nuclear transcription factors such as Elk1, activating them.**

- **Phosphorylated Elk1 joins SRF to stimulate the transcription and translation of a set of genes needed for cell division.**
<table>
<thead>
<tr>
<th>Signal Molecule</th>
<th>Site of Origin</th>
<th>Chemical Nature</th>
<th>Some Actions</th>
</tr>
</thead>
<tbody>
<tr>
<td>Hormones</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Adrenaline</td>
<td>adrenal gland</td>
<td>derivative of the amino acid tyrosine</td>
<td>increases blood pressure, heart rate, and metabolism</td>
</tr>
<tr>
<td>Cortisol</td>
<td>adrenal gland</td>
<td>steroid (derivative of cholesterol)</td>
<td>affects metabolism of proteins, carbohydrates, and lipids in most tissues</td>
</tr>
<tr>
<td>Estradiol</td>
<td>ovary</td>
<td>steroid (derivative of cholesterol)</td>
<td>induces and maintains secondary female sexual characteristics</td>
</tr>
<tr>
<td>Glucagon</td>
<td>alpha cells of pancreas</td>
<td>peptide</td>
<td>stimulates glucose synthesis, glycogen breakdown, and lipid breakdown in, e.g., liver and fat cells</td>
</tr>
<tr>
<td>Insulin</td>
<td>beta cells of pancreas</td>
<td>protein</td>
<td>stimulates glucose uptake, protein synthesis, and lipid synthesis in, e.g., liver cells</td>
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