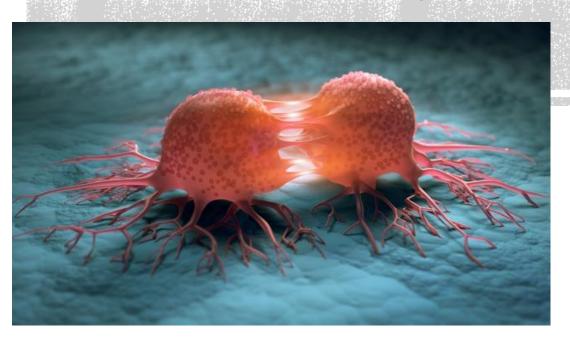
METABOLISM & CELL COMMUNICATION

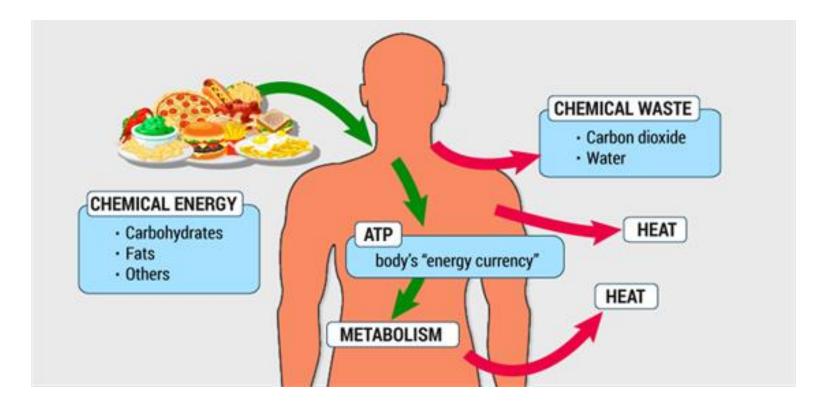


Dr. Manu Smriti Singh

Department of Biotechnology

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METABOLISM



All the chemical processes that take place in the body in order to sustain life- allowing you to breathe, pump blood, keep your brain functioning and extract energy from your food.

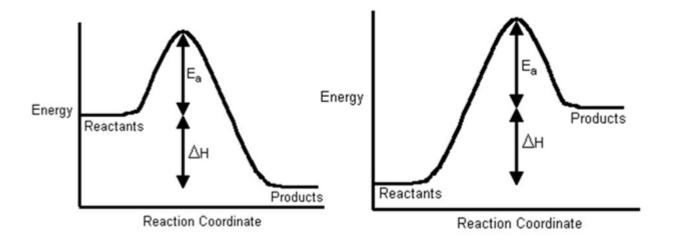
LIVING ORGANISMS AS THERMODYNAMIC SYSTEMS

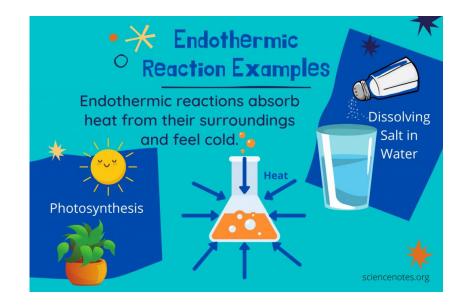
- The matter flowing into the living system contains a high energy potential.
- The matter flowing out of the system is at a low energy potential.
- The energy changes that occur between these two mass flow events are used to perform chemical and physical work processes.

- \bullet Enthalpy (ΔH)
- \Leftrightarrow Entropy (ΔS)

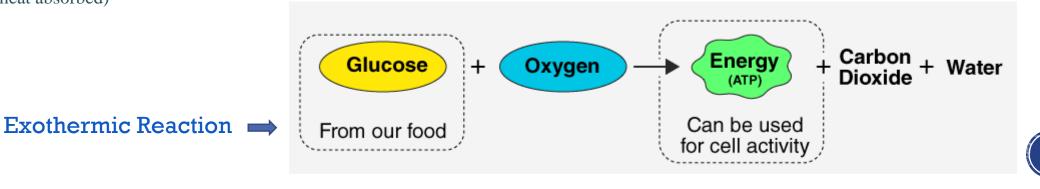
What is Enthalpy?

- ΔH, heat energy
- ENDOthermic: heat is taken in by the reactants
- EXOthermic: heat is released as a product





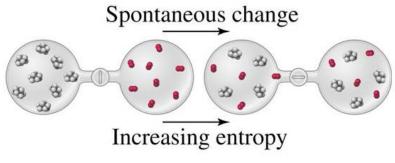
Exo (ΔH) = -ve (heat given out) Endo (ΔH) = +ve (heat absorbed)



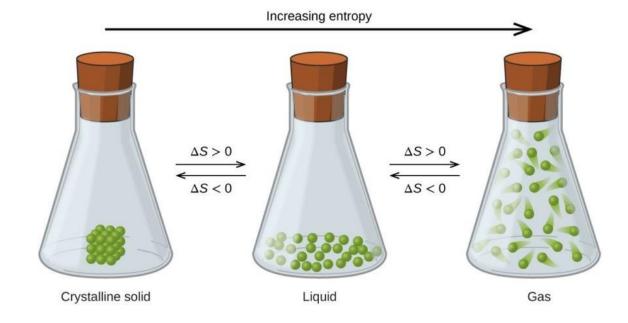
ENTROPY

Consider mixing two gases: this occurs spontaneously, and the gases form a homogeneous mixture.

There is essentially no enthalpy change involved, so why is the process spontaneous?

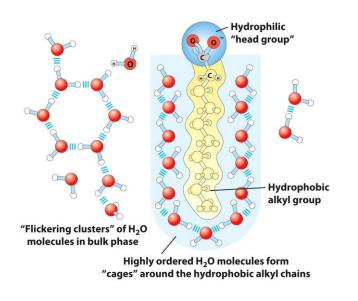


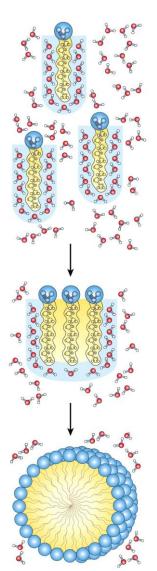
The driving force is a thermodynamic quantity called entropy, a mathematical concept that is difficult to portray visually



HYDROPHOBIC EFFECT

Suspension of a hydrophobic substance in water is thermodynamically unfavorable due to the decreased entropy of water molecules in the cage-like shell.





Dispersion of lipids in H₂O

Each lipid molecule forces surrounding H₂O molecules to become highly ordered.

Clusters of lipid molecules

Only lipid portions at the edge of the cluster force the ordering of water. Fewer H₂O molecules are ordered, and entropy is increased.

Micelles

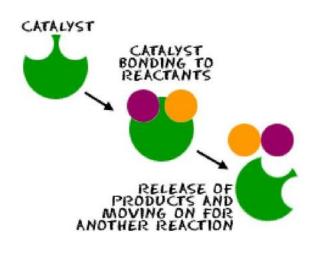
All hydrophobic groups are sequestered from water; ordered shell of H₂O molecules is minimized, and entropy is further increased.

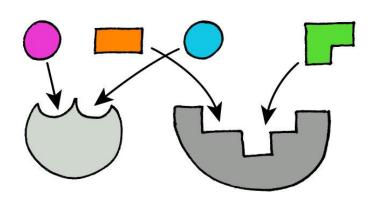
- The hydrophobic effect, and the term hydrophobic interactions, refers to the entropy-driven aggregation of nonpolar molecules in aqueous solution that occurs to minimize the ordering of water molecules with which they are in contact. This is not an attractive force, but rather a thermodynamically driven process.
- The hydrophobic effect drives the formation of membranes and contributes to the folding of proteins and the formation of double helical DNA.

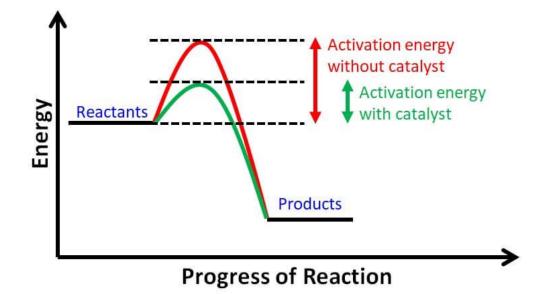


What is a catalyst?

- A compound that speeds up the rate of a chemical reaction, but is not used up in the reaction.
- Can be used over and over







A. Without a catalyst. Note the higher activation energy required for the process to start. Reaction would take longer to form and products would gradually form.

B. With the help of a catalyst, the activation energy required for the reaction goes down significantly. This means the process would go faster and yield more of the products.



Examples of Classification of Enzymes

Oxidoreductoases

oxidases - oxidize ,reductases - reduce

Transferases

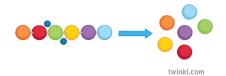
transaminases – transfer amino groups kinases – transfer phosphate groups

Hydrolases

proteases - hydrolyze peptide bonds lipases - hydrolyze lipid ester bonds

Lyases

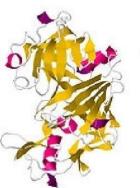
carboxylases – add CO₂ hydrolases – add H₂O



D-12 -1-1 -4

Enzyme	Reactant	Product
Pepsin	Protein	Short polypeptides
Rennin	Soluble casein (milk protein)	Insoluble casein (curdled milk)

Rennin (Chymosin)



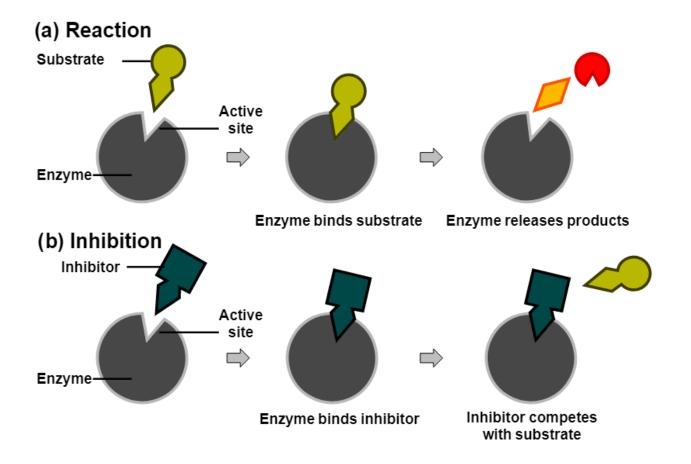
- 1. The main proteolytic enzyme found in rennet.
- 2. It is produced by Infants in the lining of the stomach.
- 3. It curdles milk, allowing a longer residence in the bowels and better absorption.

Enzyme Locations

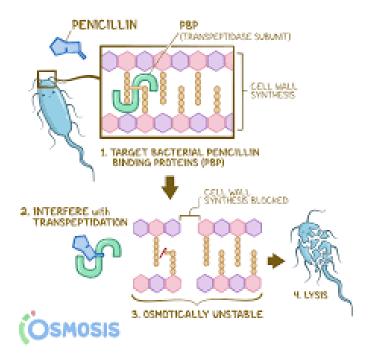
Summary of enzymes involved with digestion, what reaction they catalyse and where they are produced: -

Enzyme	Reaction Catalysed	Location Produced
Amylase	Starch → Sugars	Salivary glands; pancreas; small intestines
Protease	Proteins → Amino Acids	Stomach; pancreas; small intestines
Lipase	Lipids → Fatty Acids + Glycerol	Pancreas; small intestines

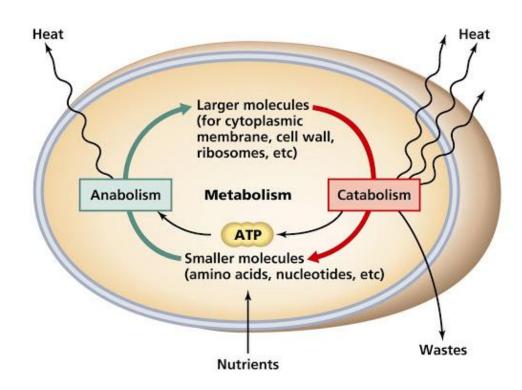
ENZYME INHIBITORS



- Penicillins inhibit a bacterial enzyme called the transpeptidase enzyme which is involved in the synthesis of the bacterial cell wall
- The β-lactam ring is involved in the mechanism of inhibition
- Penicillin becomes covalently linked to the enzyme's active site leading to irreversible inhibition



METABOLITES



The reactants, intermediates and products of metabolic pathways are referred to as metabolites

Primary Metabolites:

Enzymes, Vitamins, Lactic Acid, Proteins, Lipids, Carbohydrates

Secondary Metabolites:

Steroids, Essentials Oils, Pigments, Antibiotics, Flavonoids, Gums/Latex/ Tannins

SECONDARY METABOLITES

- Also called specialized metabolites, toxins or natural products, are organic compounds produced by <u>bacteria</u>, <u>fungi</u>, <u>or plants</u> which are not directly involved in the normal growth, development, or reproduction of the organism.
- Mediate ecological interactions, which may produce a <u>selective advantage</u> for the organism by increasing its survivability or fecundity.
- Specific within a phylogenetic group.
- Secondary metabolites often play an important <u>role in</u> <u>plant defense against herbivory and infections</u>.
- Humans use secondary metabolites as <u>medicines</u>, <u>flavorings</u>, <u>pigments</u>, <u>and recreational drugs</u>



DEFINITION AMONG **ORGANISMS** ORIGIN QUANTITY **以下在前条用的**。 PHASE OF **PRODUCTION** INVOLVEMENT IN DEFENSE REACTIONS

EXAMPLE

Primary Metabolites

Primary metabolites are compounds that are essential and directly involved in the growth, development and reproduction of an organism

Most primary metabolites are identical among most organisms

Produced during the growth phase of the cell

Produced in large quantities

The growth phase where primary metabolites are produced is sometimes called 'trophophase'

Do not participate in defense reactions

Proteins, carbohydrates, and lipids are the main primary metabolites

Secondary Metabolites

Secondary metabolites are the end products that are not directly involved in the growth, development and reproduction of an organism

Secondary metabolites are numerous and widespread

Produced during the non-growth phase of the cell

Accumulated by plant cells in very small quantities than primary metabolites

The phase during which secondary metabolites are made is called 'idiophase'

Most secondary metabolites participate in defense reactions

Alkaloids, phenolics, sterols, steroids, essential oils and lignins, are several secondary metabolites

METABOLITES IN USE

Secondary Metabolites:

Plant-Taxol/ Quinine/ Tannin/ Caffeine/ Nicotine Bacteria- Botulin Fungi- Penicillin

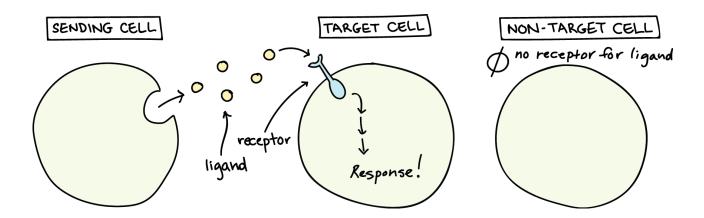
CELLS TALK IN CODES

Some UNIVERSAL PRINCIPLES of cell communication are now well known

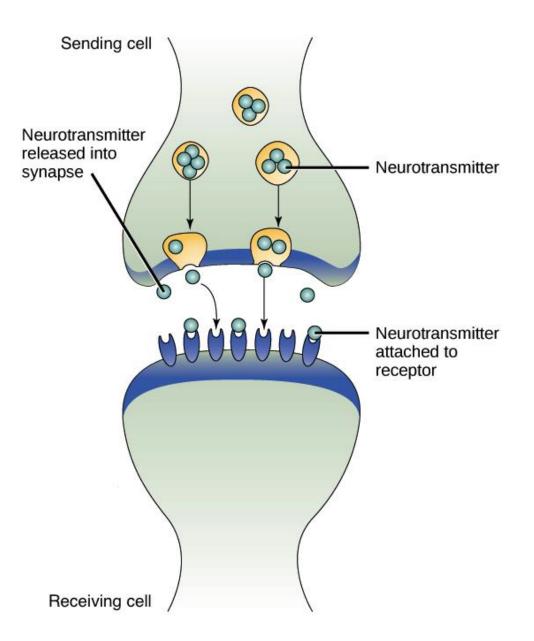
- cells may use many different signal molecules including gas (NO)
- but only a few mechanisms have survived throughout evolution.

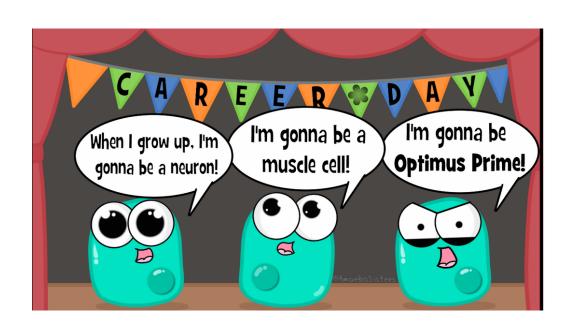
In multi-cell organisms cell-to-cell contact is critical.

- cell membranes contain specific protein-receptors, which bind & transmit extra-cellular signal molecules converting signals into specific cellular responses.



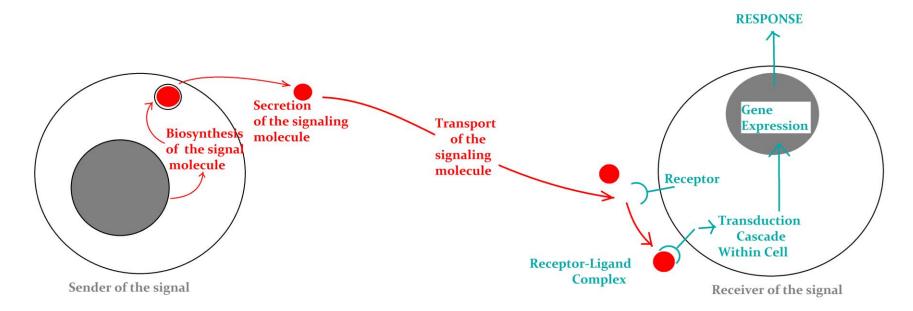
Specificity!





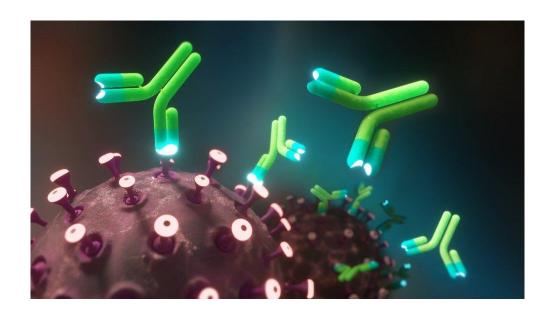
WHY DO CELLS SIGNAL?

- Regulate gene expression- (Vitamin D, Thyroid)
- Warn of possible infection
- Regulate metabolism
- Allow enzyme secretion in stomach
- Relay messages to-and-from brain-to-body and vice versa
- Fight or flight reaction



RECEPTORS

- 0.01% of the total mass of protein in a cell
- Search through:
- 1. Gene analysis
- 2. Monoclonal antibodies



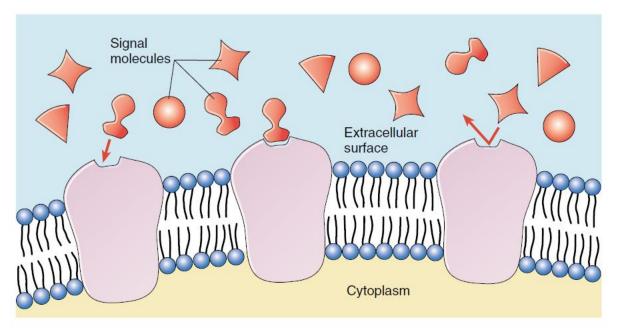
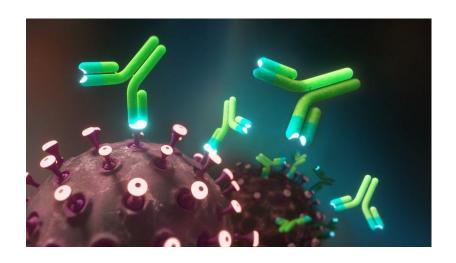
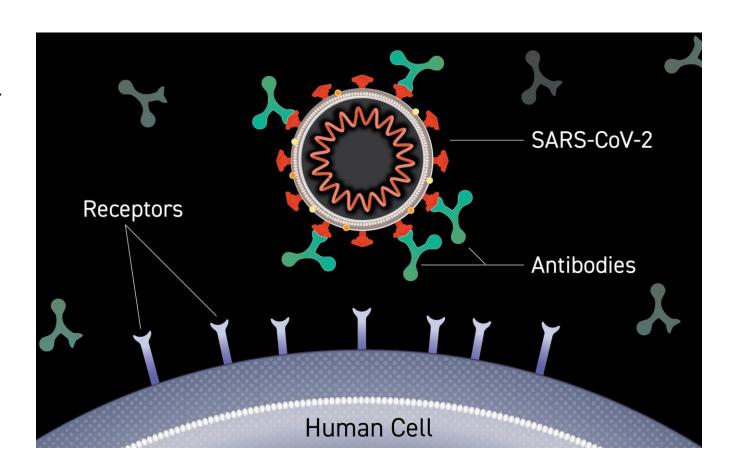


FIGURE 7.2
Cell surface receptors recognize only specific molecules. Signal molecules will bind only to those cells displaying receptor proteins with a shape into which they can fit snugly.

RECEPTORS

- 0.01% of the total mass of protein in a cell
- Search through:
- 1. Gene analysis
- 2. Monoclonal antibodies

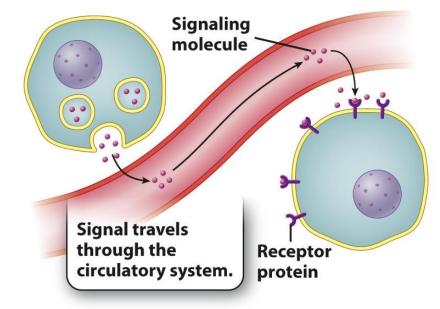




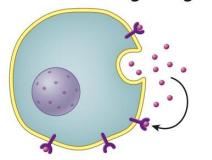
CELLS TALK IN CODES

a. Endocrine signaling

Hormones

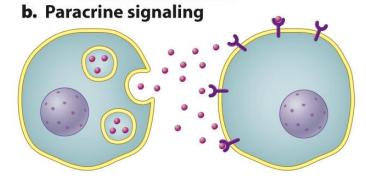


c. Autocrine signaling

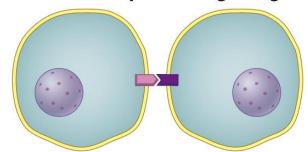


Cancer

Growth factors/ Neurotransmitters



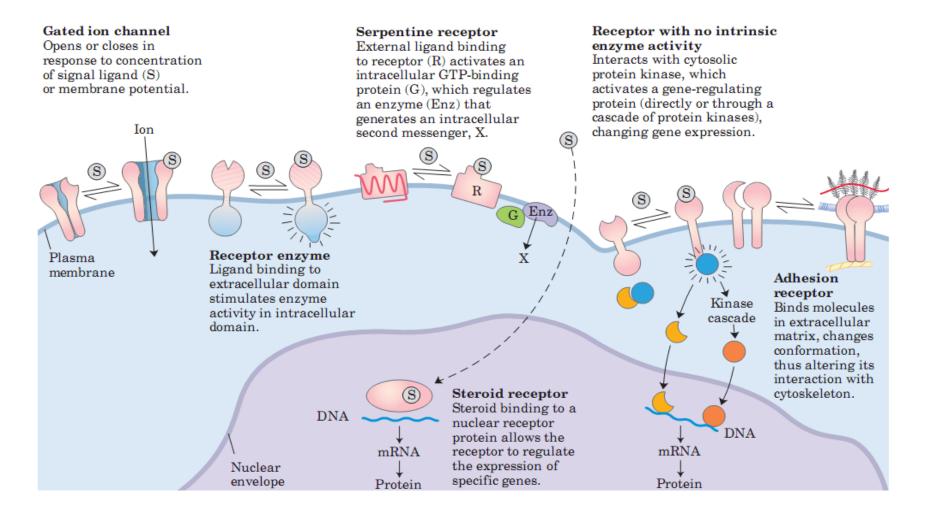
d. Contact-dependent signaling



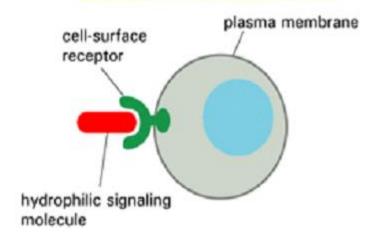
Immune cells (T-cell)

Receptors= Connect inside world with the outside world

CELL SIGNALLING



CELL-SURFACE RECEPTORS



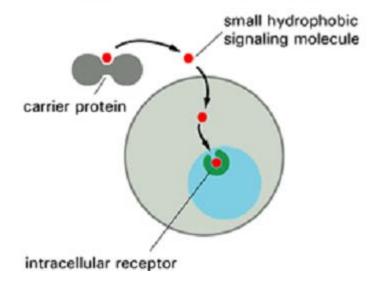
Extra cellular Hormone

- a) Hydrophilic- Glucagon, Insulin, Epinephrine and Norepinephrine
- b) Lipophilic-
 - 1) Membrane Receptor-Prostaglandins
 - 2) Nuclear Receptor- sex hormones, thyroxine, Vitamin D, Retinoic Acid

Intra cellular Hormone

- a) Proteinaceous- Kinase, Phosphatase, GTPase switch protein (ras, rab)
- b) Non-proteinaceous (sec. messenger)-cAMP, IP3, DAG, Ca2+, Phosphoinoside

INTRACELLULAR RECEPTORS





RECEPTOR COMPLEX

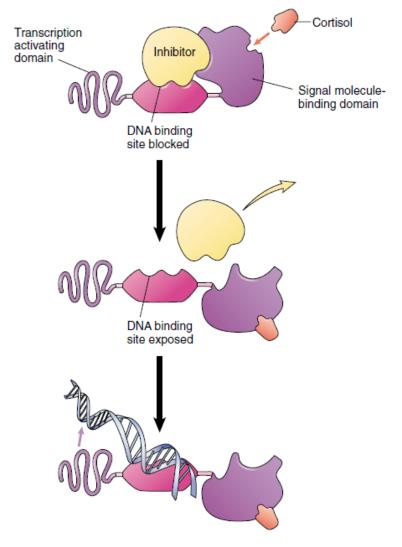
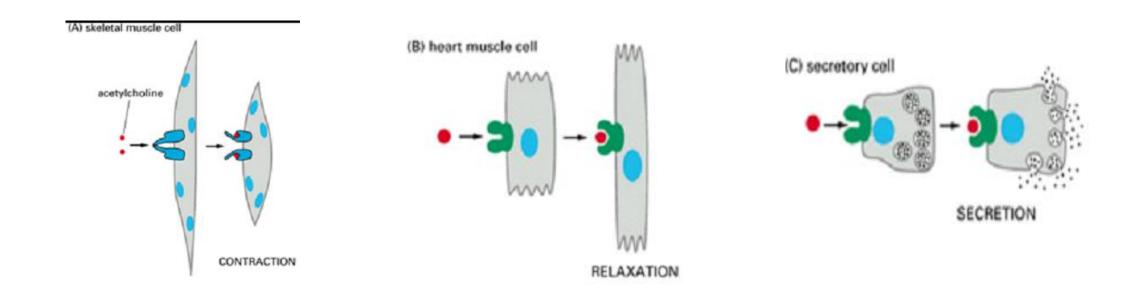


FIGURE 7.5

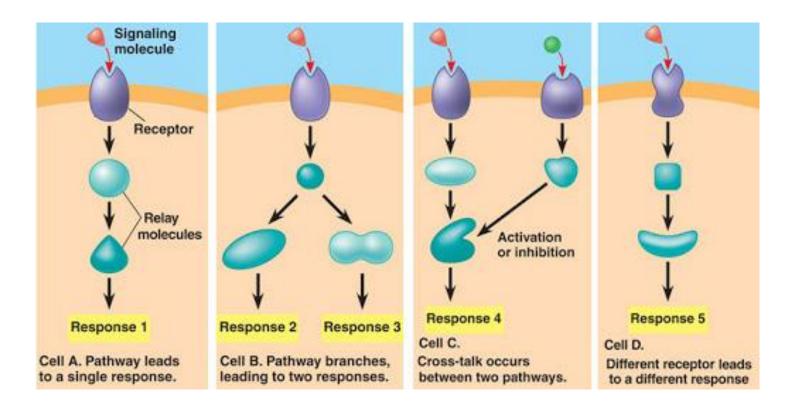
How intracellular receptors regulate gene transcription. In this model, the binding of the steroid hormone cortisol to a DNA regulatory protein causes it to alter its shape. The inhibitor is released, exposing the DNA binding site of the regulatory protein. The DNA binds to the site, positioning a specific nucleotide sequence over the transcription activating domain of the receptor and initiating transcription.

ACETYLCHOLINE AS THE SIGNALING MOLECULE



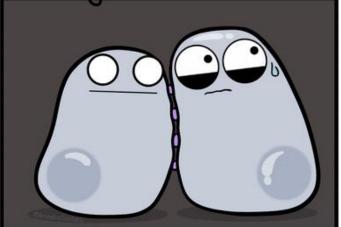
One Ligand can bind to multiple recptor types and bring about different biological response

PERMUTATION-COMBINATIONS



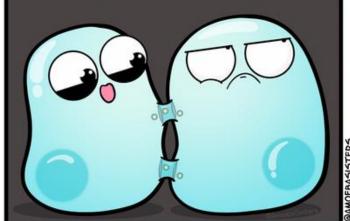
CELL JUNCTIONS

Tight Junctions



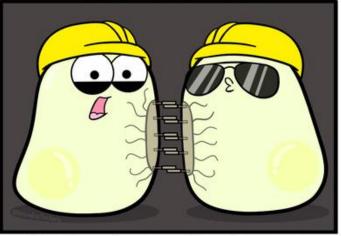
Got leaks? Not with this belt-like junction!

Gap Junctions



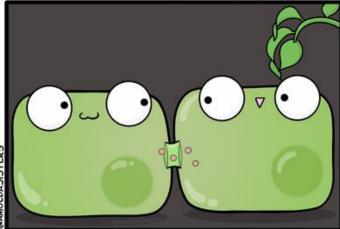
Sharing is caring! Of ions and molecules that is. [generally in animal cells]

Desmosomes



Bonds cells together for super mechanical strength!

Plasmodesmata



Similar to gap junction [but generally for plant and algae cells]

TIGHT JUNCTION VERSUS GAP JUNCTION

TOTT	IUNCTION	

A specialized connection of two adjacent animal cell membranes, such that, space usually lying between them is absent

Also known as occluding junctions and zonulae occludentes

Occur in the epithelia of vertebrates

Contain plasma membranes, which are very tightly pressed against each other due to bound proteins

Contain proteins called claudins

Regulate the movement of water and solutes between epithelial layers

Prevent the leakage of extracellular fluid across the epithelial cell layer

GAP JUNCTION

A linkage of two adjacent cells consisting of a system of channels extending across a gap from one cell to the other, allowing the passage

Also known as nexus and macula communicans

Occur in all types of tissues, except in fullydeveloped skeletal muscles and mobile cells types

Consist of cytoplasmic channels from one to the other cytoplasm, surrounding with membrane proteins

Contain proteins called connexins

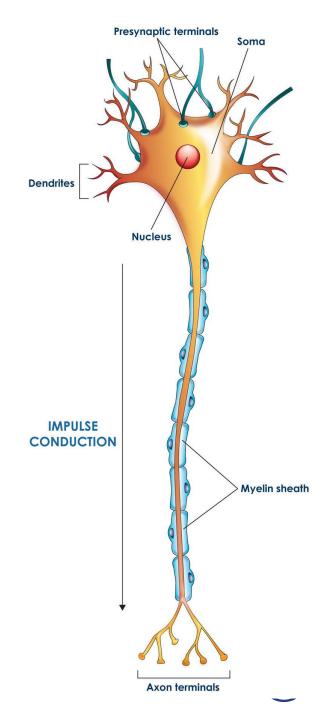
Allows the direct chemical communication between adjacent cytoplasm

Allow the movement of ions. sugars, amino acids, and other small molecules between cells

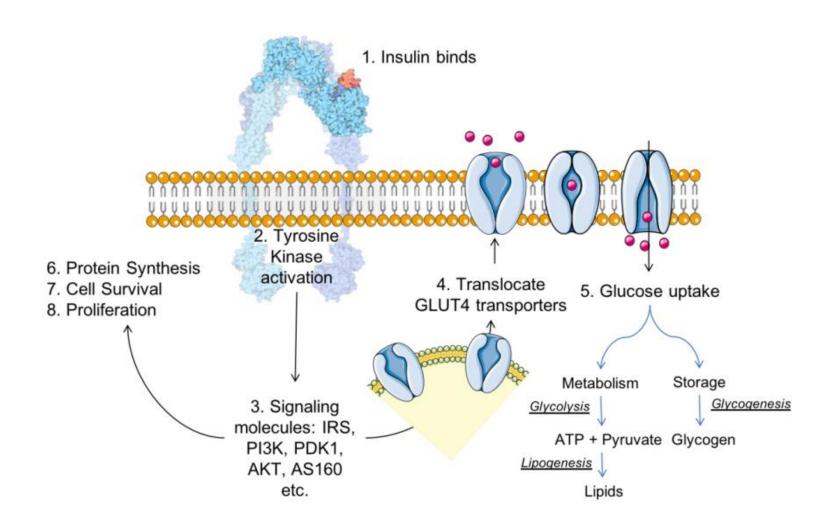
Visit www.PEDIAA.com

Multiple Sclerosis

Myelin sheath that protects nerve cells disappears Brain and spinal cord Signal do not pass

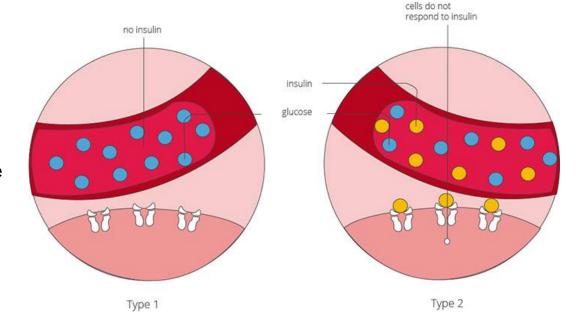


INSULIN SIGNALLING PATHWAY:



Diabetes

Target cell receptor not responding to Insulin signal Diabetes 1- insulin signal is unable to be produced Diabetes 2- cells have lost the ability to respond to the signals, resulting in abnormally high and dangerous sugar levels in the blood



Brain Stroke

- Abnormally high amount of glutamate secreted by dying brain cells
- Kills healthy brain cells (excitotoxicity)
- Extensive brain damage



Cancer

Breakdown of multiple signaling pathways Uncontrolled proliferation of cells- cancer

