# **PHSI2005**

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# **Concepts in Physiology**

### Homeostasis

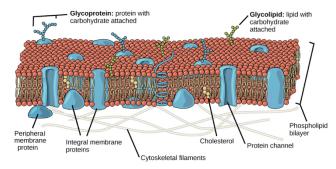
- A set of steady states in which an organism operates comfortably
- Constant energy input and losses to maintain

## The plasma membrane

- Cytoskeletal filaments allow for stability, contraction and relaxation, motility and shape change
  - Example: RBC squish to pump the blood around (reduces viscosity)
- Transmembrane proteins allow selectively entry of molecules that can't pass through the membrane normally

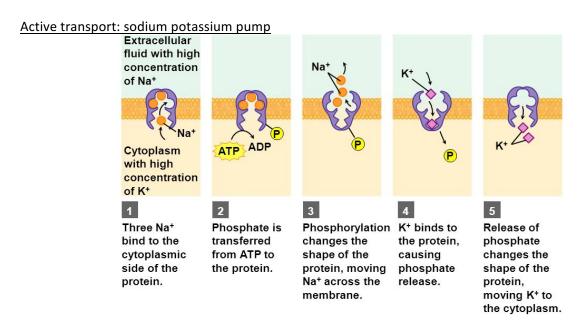
Receptor proteins receive and interpret

- signals
  Specialised lipids alter permeability to small, hydrophobic molecules
- Permeable to small, hydrophobic molecules
- Barrier to large hydrophobic molecules, atomic ions, charged molecules, polar molecules



### Transport

- Diffusion = spread of molecules from areas of higher concentration to areas of lower concentration
  - Simple: straight through the plasma membrane
  - Carrier-mediated: substrate binding causes shape change in transport so molecule can move through
    - (1) Uniport: one kind of substrate
    - (2) Symport: two plus substrates in same direction
    - (3) Antiport: substances in opposite directions
  - Channel-mediated: molecules move through a channel protein
- Osmosis = molecules of a solvent pass through a semipermeable membrane from a less concentrated solution into a more concentrated one
  - Through a specific channel protein (aquaporin) or through lipid bilayer (much slower)



#### Facilitated diffusion of glucose

- Glucose is brought into the cell down its concentration using GLUT transporter
- Diffusion reaches equilibrium when concentration inside = concentration outside
- Conversion of imported glucose into G6P keeps intracellular glucose concentration low so that diffusion never reaches equilibrium

### RBC's in solution

- Isotonic: concentration same inside and outside RBC
- Hyperosmostic: low solutes inside the cell, water moves out and cell shrivels
- Hypoosmotic: low solutes outside the cell, water rushes in and the cell swells and bursts

# Secondary transport: SGLT transporter

- Na binds to carrier
- Na binding creates high affinity for glucose
- Glucose binding changes carrier conformation so binding sites now face ICF
- Na released into cytosol where [Na] is low
- Release changes glucose binding site to low affinity and glucose is released

# Negative feedback: feedback used to decrease size of input

- (1) Stimulus produces change
- (2) Receptor detects change
- (3) Input information sent (afferent) to control centre
- (4) Output information sent (efferent) to effector
- (5) Response of effector feeds back to reduce the effect of stimulus and returns variable to homeostatic level

# <u>Positive feedforward: to change a variable in anticipation of a process</u>

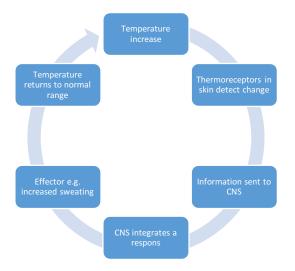
Anticipation of eating food so we salivate more

# Positive feedback: feedback used to increase size of input

- Baby drops lower in uterus to initiate labour
- Cervical stretch
- Stimulates oxytocin release
- Causes uterine contractions
- Push baby against cervix
- More cervical stretch

### Chemical signalling

- Needs a signal input (ligand) to bind to a receptor
- This changes conformation, amplifying the signal
- Cell takes this information and decides how to respond
- Ligand-receptor interactions
  - Signal molecule acts as a ligand, binding with specific proteins on the membrane
  - This induces a change in shape of a receptor protein, inducing a cellular response
- Signal amplification



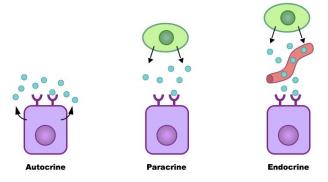
- Receptor activates stage 1 protein which activates stage 2 protein ... stage 3
- Stimulation of a cascade of protein kinases to amplify the signal produces a strong response inside the cell
- Phosphorylation
  - Signal transduction pathways involve phosphorylation cascades series of kinases successively add phosphate groups to next one in line, activating them
- Signal transduction converts cell signal on cellular surface into a specific response

## Emergent properties of the cell

- Type of cell
- State of cell
- Type of ligand
- Concentration of ligand
- Receptor and receptor subtype
- Receptor concentration
- Number of different ligands

### Types of signalling

- Endocrine: cells respond to factors produced by distant cells (hormones)
- Paracrine: cells respond to secretion of nearby cells
- Autocrine: cells have receptors for their own secreted factors



### Types of membrane receptors

- Receptor channel: ligand binds to open or close channel
- Receptor-enzyme: ligand binds to a receptor and enzyme activates an intracellular enzyme
- G protein coupled receptor: ligand binds to receptor, G protein coupled receptor alters the enzyme activity or opens an ion channel
- Integrin receptor: ligand binds to integrin receptor to alter the cytoskeleton

### Oxygen transport and use

- Oxygen binds to haemoglobin
- Oxygen is small, uncharged and non-polar easily crosses cell membranes, diffuses rapidly
- High oxygen partial pressure in lungs
- 2% oxygen dissolved in plasma, 98% attached to haemoglobin
- Low oxygen partial pressure at tissues (increases concentration gradient)