

STUDY GUIDE:

B I O L O G Y **HL**

www.ib.academy



IB Academy Biology Study Guide

Available on learn.ib.academy

Authors: Lea Knežević, Paulina Núñez, Matthew Hugh Tan
Contributing authors: Annika Semmler, Laetitia Boon, Elisa Miranda

Design

Rational

Typesetting



*This work may be shared digitally and in printed form,
but it may not be changed and then redistributed in any form.*

Copyright © 2019, IB Academy
Version: BioHL.3.0.191218



This work is published under the Creative Commons
BY-NC-ND 4.0 International License. To view a copy of this
license, visit creativecommons.org/licenses/by-nc-nd/4.0

This work may not be used for commercial purposes other than by IB Academy, or parties directly licenced by IB Academy. If you acquired this guide by paying for it, or if you have received this guide as part of a paid service or product, directly or indirectly, we kindly ask that you contact us immediately.

Laan van Puntenburg 2a
3511ER, Utrecht
The Netherlands

ib.academy
info@ib.academy
+31 (0) 30 4300 430

INTRODUCTION

Welcome to the IB.Academy Study Guide for Biology High Level.

We are proud to present our study guides and hope that you will find them helpful. They are the result of a collaborative undertaking between our tutors, students and teachers from schools across the globe. Our mission is to create the most simple yet comprehensive guides accessible to IB students and teachers worldwide. We are firm believers in the open education movement, which advocates for transparency and accessibility of academic material. As a result, we embarked on this journey to create these study guides that will be continuously reviewed and improved. Should you have any comments, feel free to contact us.

In this Biology HL guide we present everything that is vital for your final exam. Each section is clearly marked relating to the topics of the syllabus. The material is summarised in short texts that include various definitions, images, examples and labeled diagrams which will help you with your studies. This study guide also contains useful tips and info boxes that will assist you along the way, telling you which components are important for your exams.

For more information and details on our revision courses, be sure to visit our website at ib.academy. We hope that you will enjoy our guides and best of luck with your studies.

IB.Academy Team

TABLE OF CONTENTS











	1. Cell biology	7
	2. Molecular biology	31
	3. Genetics	59
	4. Ecology	93
	5. Evolution and biodiversity	105
	6. Human physiology	119
	7. Nucleic acids	153
	8. Metabolism, cell respiration and photosynthesis	161
	9. Plant biology	177
	10. Animal physiology	193

TABLE OF CONTENTS

CELL BIOLOGY

1

1.1. Cell theory

8

- Sizes of cells
- Cell properties
- Examples of stem cell use

1.2. Cells and membrane transport

15

- Eukaryotic and prokaryotic cells
- Cell membrane
- Membrane transport
- Osmolarity

1.3. Origin of cells

24

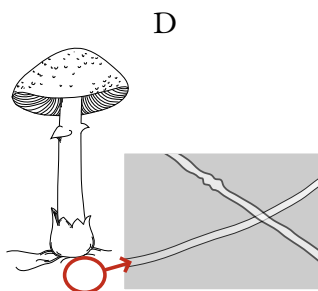
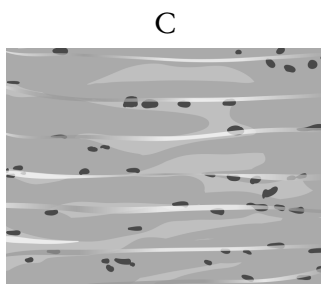
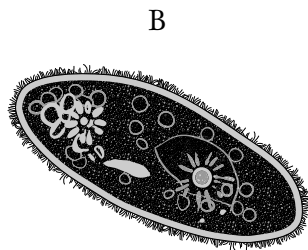
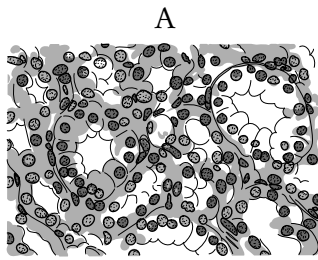
- Pasteur's soup
- Formation of organic molecules
- Endosymbiotic theory

1.4. Cell division

25

- Cell cycle
- Phases of mitosis

1.1 Cell theory



1. **All organisms are composed of cells:**
microscopic examination of many organisms has shown that they are all composed of cells; unicellular organisms are still composed of one cell that performs all functions of life.
2. **Cell is the basic unit of life:**
cell as a whole can perform the functions of life, while its individual components cannot.
3. **All cells originate from a pre-existing cell:**
spontaneous generation of cells is not possible; a cell needs to divide to create another cell.

Even though most organisms fit well into the first two points of the cell theory (A and B), some organisms and tissues seem to contradict it. Muscle fibres are fused, elongated cells with multiple nuclei and as such differ from the common definition of a cell (C). Similarly, fungal hyphae often don't contain dividing walls and are made up multiple fused thread-like cells (D).



7 Functions of life carried out by cells

- 1 Nutrition
- 2 Metabolism
- 3 Excretion
- 4 Response
- 5 Homeostasis
- 6 Growth
- 7 Reproduction

If these are hard to remember, think about the following:

Every living being has to eat^①, process the food it has eaten^② and excrete the waste^③, take in the signals from the environment and respond to them^④, accordingly make sure that it is well balanced^⑤ (food-wise, heat wise, etc.) and finally, use all that to grow^⑥ and pass on its genes^⑦.

Example.

Example: Paramecium is a unicellular organism widely used as an example of a functional unit of life.

The Paramecium:

- Is surrounded by cilia that allows it to move. (*Response*)
- Engulfs food via a specialized membranous feeding groove called a cytostome. (*Nutrition*)
- Encloses food particles in small vacuoles with enzymes for digestion. (*Metabolism*)
- Removes solid waste via an anal pore and liquid wastes are pumped out through contractile vacuoles. (*Excretion*)
- Allows essential gases to enter and exit the cell via diffusion. (*Homeostasis*)
- Grows in size and divides asexually (fission) (*Reproduction and growth*)

1.1.1 Sizes of cells

Surface area to volume ratio

Surface Area: of a cell is related to the rate of exchange of materials or how fast/slow a cell takes in food/gasses and excretes wastes products.

- The higher the surface area, the higher the exchange rate as there is more physical membrane where exchange can happen.

Volume: of a cell is related to the rate of metabolic reactions occurring in a cell or how much nutrient processing and waste production is occurring in a cell at a given time.

- The higher the volume of a cell, the higher the metabolic rate as the more nutrients are needed/used and thus more waste produced.

Cell growth is limited by two features of the cell: *surface area* and *volume*.

When the cell volume increases, the surface area increases comparably less. This limits the size of a cell because:

- the cell must be able to transport enough food/waste through the surface
- compared to the food needs/excrement production, which is determined by the cell volume.

This is linked to cell division, as following a period of growth a cell will divide in order to increase the surface area to volume ratio and function more effectively. In addition, some cells increase their surface area to volume ratio by creating folds in the plasma membrane, which creates more surface area to cope with a large exchange of materials. This can be seen in the cells of the intestinal lining.

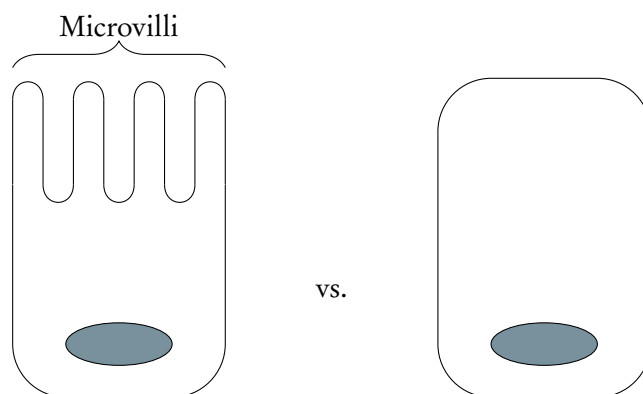
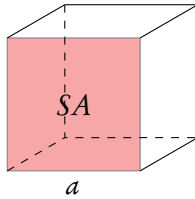


Figure 1.1: Microvilli increases surface area for exchange.

Example.

Example surface area (SA) and volume (V) calculation



Imagine a cell as a cube with sides of length a . The surface area (SA) can be calculated by adding the SAs of the six faces of the cube and the volume (V) by multiplying the sides.

$$SA = 6 \times a^2 \quad \text{and} \quad V = a^3$$

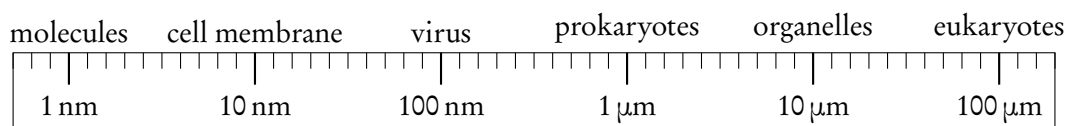
Now, compare the growth of cells, starting with $a = 1$, $a = 5$ and $a = 10$.

	size of 1	size of 5	size of 10
side = a	$a = 1$	$a = 5$	$a = 10$
$SA = 6 \times a^2$	$SA = 1$	$SA = 150$	$SA = 600$
$V = a^3$	$V = 1$	$V = 125$	$V = 1000$

Up to a certain size the SA still exceeds the V, and the cell would be able to import and export enough materials to sustain its life. But as the cell grows bigger the volume will exceed the SA, at which point the cell cannot transport enough materials in and out of the cell to keep up with its food needs/waste production.

Typical cell sizes

Cells have different sizes, from one organism to another as well as within an organism. This difference arises from the different cell functions and needs. The following scheme should help you compare the sizes of different cells:



Calculating magnification

Most cells are invisible to the naked eye therefore we use microscope magnification to be able to see them.



$$\text{Image of the cell} = \text{real cell size} \times \text{magnification}$$

Example.

Real cell size

Exam question: *Image of a cell, measurable by a ruler. Magnification in the corner.*

Calculate the real size of the cell in the picture or the magnification factor

Solution:

$$\text{Image of the cell} = \text{real cell size} \times \text{magnification}$$

therefore

$$\text{Real cell size} = \frac{\text{Image of the cell}}{\text{magnification}}$$

Numbers worked out.

Remember to always use the same units

- 1 mm = 1000 micrometers (μm)

Microscopes

An electron microscope has a far greater resolving power than a conventional light microscope, meaning an electron microscope can be used to create images of smaller objects with greater resolution.

The limit of resolution is determined by the wavelength of the incident light/electrons. And since the wavelength of electrons is much smaller than that of visible light, they can be used to view objects much smaller and in much more detail.

1.1.2 Cell properties



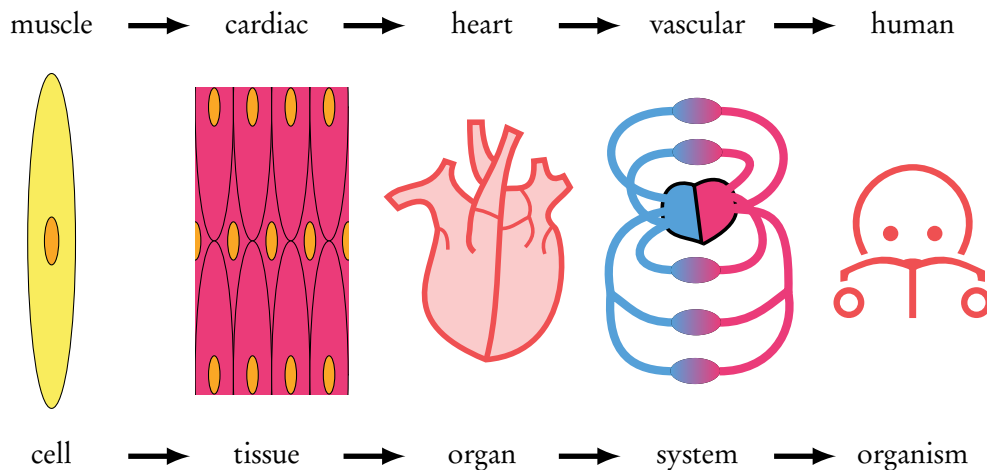
In multicellular organisms, each cell has its own function and cooperates with other cells to form an organism.

Emergent properties

Emergent properties are properties that emerge from the interaction of the individual cell components creating new functions. Multicellular organisms can realize a number of complex functions that individual cells cannot perform on their own by collaborating with one another.

Example.

You can easily visualize this by thinking about how individual cardiac cells form tissues that work together to create an organ (the heart). Then the heart together with tissues that formed vessels can function as the vascular system, transporting blood across the body. The latter cannot be done by one single cell, but is possible by the collective actions of many cells together. Finally, different systems can create an organism, like a human, who can also perform several functions that a single cell could not do on its own.



Cell differentiation

Although all cells possess the same genes, expressed genes are the ones that are “switched on” while all other genes are “switched off”. Cell differentiation is the process whereby different genes are put on “lockdown” to reach a very specific cell type with particular functions brought by the available active genes.

Stem cells

Undifferentiated cells that can divide and have the capacity to differentiate into any cell type and thus acquire any function. Therapeutic sources of stem cells include umbilical cord blood, bone marrow and human embryonic stem cells

1.1.3 Examples of stem cell use

Example.

Stadgardt's disease

Stadgardt's disease is a degenerative disease of the eye (retinal cells) leading to blindness. Human embryonic stem cells are obtained from unsuccessful in vitro fertilizations. These cells are differentiated in the lab towards retinal cells and injected into the eye of patients. The new cells replace the degenerate cells in the retina and restore vision

Example.

Leukaemia

Leukaemia is the cancer of white blood cells (immune cells). Human cord blood is collected after childbirth. The cord blood contains stem cells that differentiate into white blood cells. A patient with leukaemia is irradiated and given chemotherapy to kill all cancerous white blood cells. The killed cells are then replaced by the matching cord blood cells which are able to differentiate into all kinds of white blood cells in the patient

1.2 Cells and membrane transport

1.2.1 Eukaryotic and prokaryotic cells

The main distinctions between the eukaryotic and prokaryotic cells relate to their size and complexity. Prokaryotes, also known as bacteria, are unicellular organisms with a simple, non-compartmentalized structure. Eukaryotes, which can be unicellular or multicellular organisms, are usually bigger with a complex organelle based function and compartmentalization.

Compartmentalization means to have structures wrapped around a membrane, creating different discrete compartments. This allows regions in the cell to have specific functions and environments.

Example.

Lysosomes

Lysosomes are membrane bound organelles responsible for the digestion of materials. In order to break down materials they need to have a very acidic pH of around 4.5–5. This pH is achieved with hydrogen pumps in the organelle's membrane that pump protons inside the organelle. Without compartmentalization, this specific concentrated acidic environment would not be possible, and would disrupt normal cell functioning.

Prokaryotic cells

Make sure you can draw and clearly label this yourself!

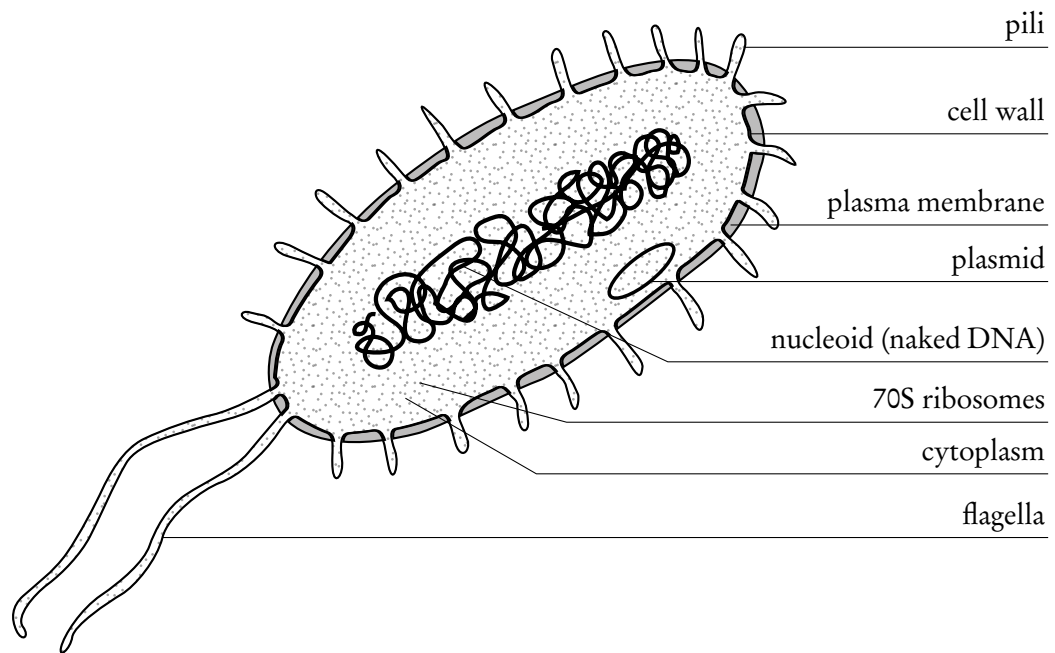


Figure 1.2: Prokaryotic cell.

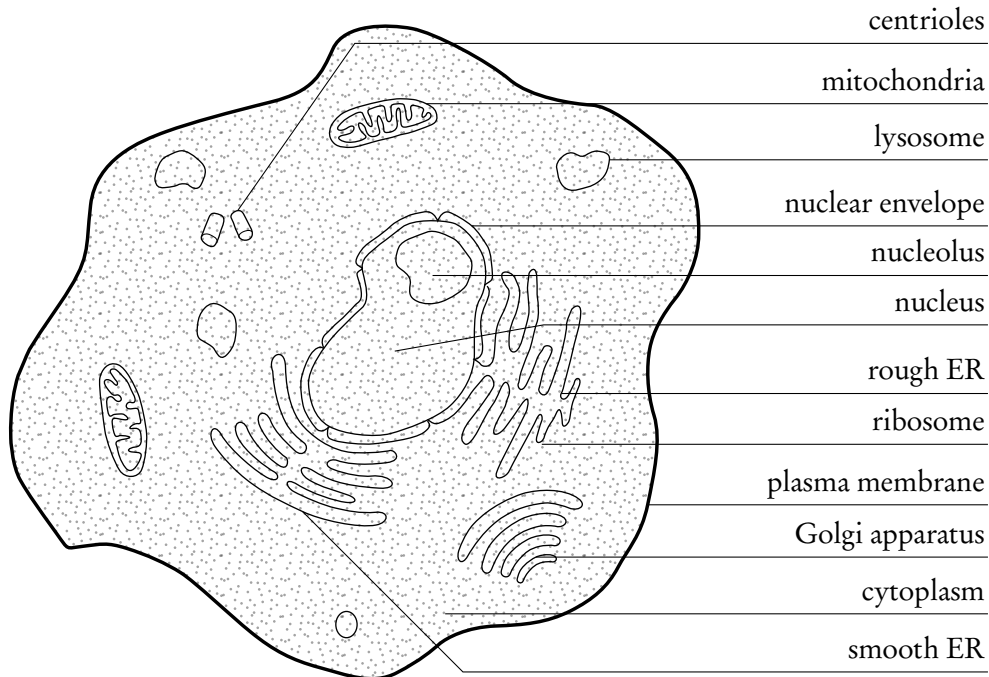
The bold structures in the table are shared between the prokaryotic and eukaryotic cells.

Table 1.1: Functions of prokaryotic cells.

Structure	Function
Capsule	Protection
Cell wall	Protection and pressure maintenance
Cell membrane	Transport of materials
Cytoplasm	Contains enzymes, food..., medium for cellular processes
Ribosomes	Protein synthesis
Nucleoid	DNA containing area not enclosed by a membrane
Plasmid	Extra genetic material (e.g., antibiotic resistance genes)
Pilli	Communication, DNA exchange, attachment
Flagellum	Movement

Eukaryotic cells

eu = good/true
karyon = nucleus } → organisms with a nucleus



Make sure you can draw and label this yourself!

Figure 1.3: Eukaryotic Cell.

There are two types of eukaryotic cells: pancreatic cell (animal) and mesophyll cell (plant).

Table 1.2: Comparison of animal cell and plant cell.

Structure	Function	Animal cell	Plant cell
Ribosome	Protein synthesis	✓	✓
Rough endoplasmic reticulum	Protein modifications	✓	✓
Golgi apparatus	Protein packaging	✓	✓
* Mitochondrion	Site of cell respiration	✓	✓
* Nucleus	Contains chromosomes (DNA)	✓	✓
Lysosome	Degradation enzyme storage	✓	✓
Centrioles	Chromosome separation during mitosis	✓	✗
Vacuole	Food and water storage	✓	✓
Cell Wall	Maintenance of cell pressure	✗	✓
* Chloroplast	Site of Photosynthesis (food production)	✗	✓
* Centrosomes	Contain microtubules, move to opposite poles during mitotic division	✓	✓

* indicates double membrane bounded organelles.

Comparison of eukaryotic cells and prokaryotic cells

Table 1.3: Comparison of prokaryotic cells and eukaryotic cells.

Prokaryotic Cell	Eukaryotic Cell
Naked DNA	DNA wrapped around
DNA in nucleoid	DNA enclosed by a nuclear region
DNA circular	DNA linear
No membrane bound structures	Membrane bound structures such as mitochondria, ER, Golgi apparatus present which compartmentalize functions
Plasmids present	No plasmids
Mitochondria not present	Mitochondria always present
Ribosomes smaller (70S)	Ribosomes larger (80S)

Remember that in a **compare and contrast** question in the exam, you only get a point for differences if you give the alternative for each thing being compared. So it is not enough to say Prokaryotes have DNA at the nucleoid region, but you must also mention that Eukaryotes have DNA enclosed in the nuclear membrane.

Besides in their structure, the two types of cells also differ in their mode of division. Eukaryotic cells divide by **mitosis** (discussed later) while prokaryotic cells divide by **binary fission**.

1.2.2 Cell membrane



Cell membrane Cell membranes are an assembly of different components that encloses cells. This includes phospholipids, cholesterol, proteins, and lipoproteins.

Phospholipids molecules composed of a glycerol head with a negatively charged phosphate group and two hydrocarbon lipid tails. This makes phospholipids amphipathic, meaning that they have two opposite properties:

- Their heads are hydrophilic (water loving) making them polar.
- Their tails are hydrophobic (water hating) making them non-polar.

Make sure you can draw and label a phospholipid diagram (Figure 1.4).

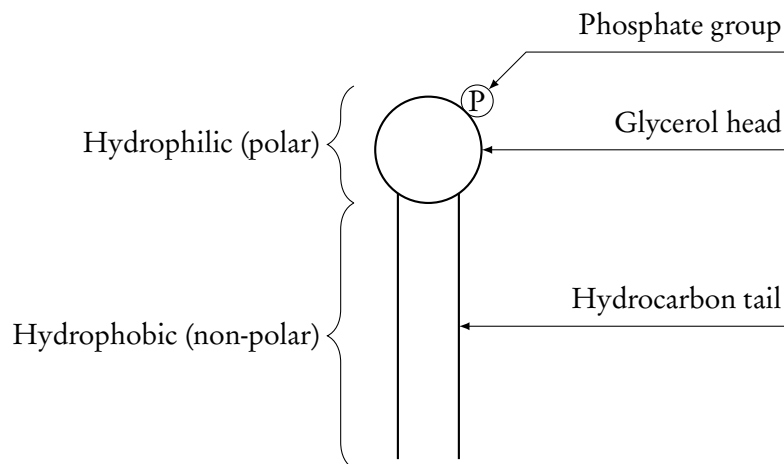


Figure 1.4: Phospholipid diagram.

This will usually give you an extra point in essay questions concerning the plasma membrane.

Notice the head / tail structure of the phospholipid: their amphipathic properties cause them to spontaneously arrange into a bilayer in an aqueous environment. This is due to the fact that water is polar, and therefore the hydrophobic, non-polar portion of the phospholipids will want to be shielded from water by the hydrophilic, polar heads of the phospholipids.

Due to these interactions, the plasma membrane is very stable but is said to be **fluid**. This means that the tails will always be facing tails, and the heads will always face outside, but the position of individual phospholipids in a layer may change.

This property of the membrane also allows it to host a variety of other molecules, like proteins and cholesterol. This makes it look like a **mosaic**.

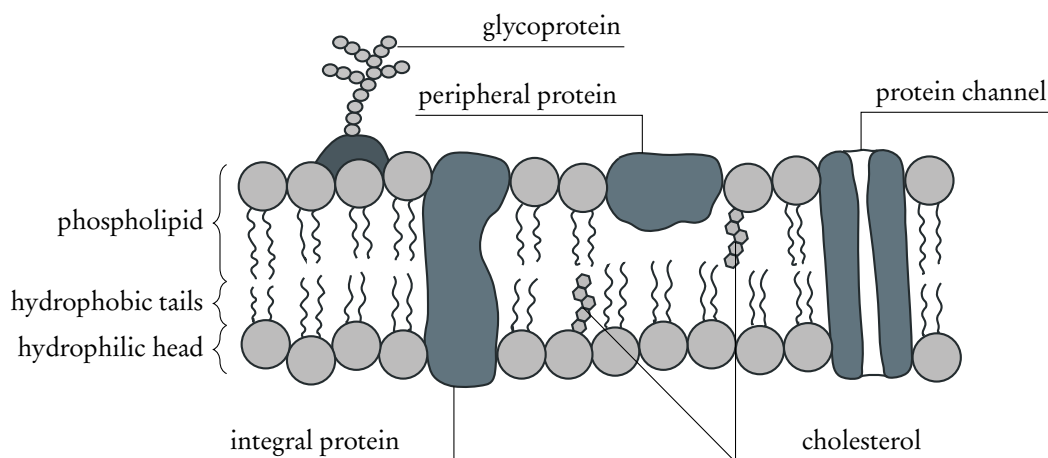


Figure 1.5: Phospholipid molecules form a phospholipid bilayer, which together with proteins and cholesterol forms cell membranes.

Cholesterol keeps the fluidity of the membrane constant at a variety of temperatures. When it is cold, it increases fluidity and when hot, it makes the membrane more rigid. This is important to maintain a constant environment for cellular processes to occur.



Integral Proteins span the lipid bilayer and are permanently attached to it by polar interactions with the phospholipid heads and nonpolar interactions with the tails. These can be for example receptors or transport proteins.

Peripheral Proteins are attached to the inner or outer side of the membrane by non-covalent interactions with the surface of the membrane or other integral proteins. These are for example enzymes attached to the outside of cells.

Membrane proteins fulfill various functions:

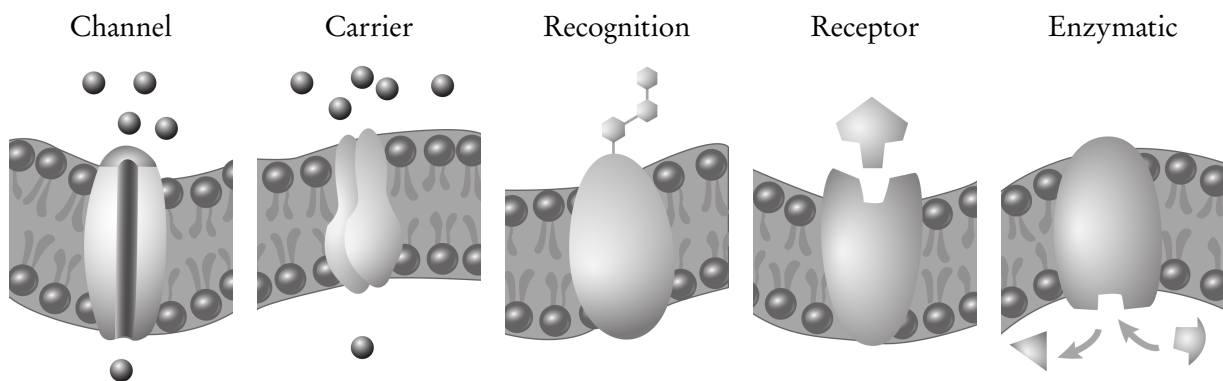


Figure 1.6: Phospholipid molecules form a phospholipid bilayer, which together with proteins and cholesterol forms cell membranes.

1.2.3 Membrane transport

The lipid bilayer membrane is semi-permeable and selective, which means that

Semi-permeable: only certain molecules (small, polar) can freely cross the membrane

Selective: with the use of transport proteins, it can select what comes in and out and what does not

Recall that the main function of the plasma membrane is transport. Generally, transport is defined as passive or active.



Active transport movement of molecules from an area of lower concentration to an area of higher concentration, with the use of energy (against the concentration gradient)

Passive transport movement of molecules from an area of higher concentration to an area of lower concentration (down the concentration gradient)

Passive transport is further divided into two types of diffusion, simple and facilitated.

Simple diffusion passive transport of molecules through a membrane, without the need of protein channels (oxygen diffusion)

Facilitated diffusion passive transport of molecules **facilitated** by channel or carrier proteins (sodium transport, calcium transport)

Osmosis is a form of passive transport that only refers to the movement of water. The water, like other particles in passive transport, moves from the area where there is more of it, to the area where there is less of it. However, osmosis is defined in terms of the concentration of dissolved molecules:



Osmosis movement of water from the area of low solute concentration to the area of high solute concentration.

Active transport, like (passive) facilitated diffusion requires proteins. However, these proteins use energy in form of ATP to pump molecules against their concentration gradient. There are two types of active transport:

Primary: direct use of metabolic energy for transport of molecules against concentration gradient.

Secondary: coupling the movement of one molecule against the concentration gradient with the movement of another along the concentration gradient of the second molecule, often created by primary active transport.

Sodium-potassium pump is such a protein, and can be found in many cells including neurons. This pump is described in more detail in the “Human physiology” chapter, but now consider the following points:

- sodium potassium pump is an integral protein that uses ATP to transport molecules across a membrane;
- it transports sodium out of the cell, and potassium into the cell;
- it works against the concentration gradients of both sodium and potassium;
- for every three sodium molecules it transports out, two potassium molecules are transported in.

You should consider two more types of active transport that involve vesicle transport, rather than protein pumps.



Exocytosis Exocytosis: Transport of molecules in secretory vesicles that fuse with the plasma membrane upon contact to release the contents outside of the cell

Endocytosis transport of molecules into the cell through invagination of the plasma membrane and formation of the phospholipid vesicle containing the molecule.

1.2.4 Osmolarity



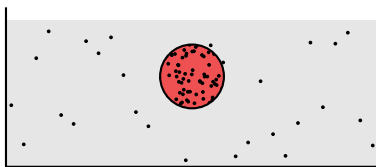
Osmolarity Is a measure of solute concentration (osmol/L) in a given system. A system can be a petri dish, a cell, an organism, etc.

Hypotonic low osmolarity, or low solute concentration, meaning that the system loses water

Hypertonic high osmolarity, or high solute concentration, meaning that the system will gain water

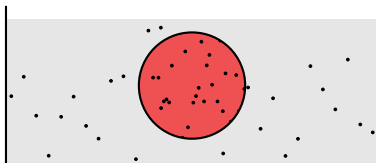
Imagine a potato cube in a water bath. Mind you that the potato has a much smaller volume compared to that of the water tank.

Hypotonic



The ratio of solutes to water inside the potato is much higher than that same ratio in a water bath.

The bath is *hypotonic* compared to the potato.

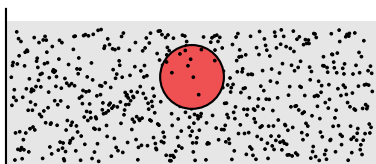


The water moves from the bath into the potato, making the potato swell.

The ratio of water to solute in the potato is the same as the one in the bath.

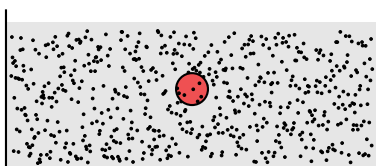
Recall that in osmosis, the water moves from where there is more of it to where there is less of it

Hypertonic



The water bath is more saturated with its solute compared to the potato.

The bath is a *hypertonic* solution compared to the potato.



The water inside the potato will pass into the water bath, trying to dilute it to the same concentration as in the potato.

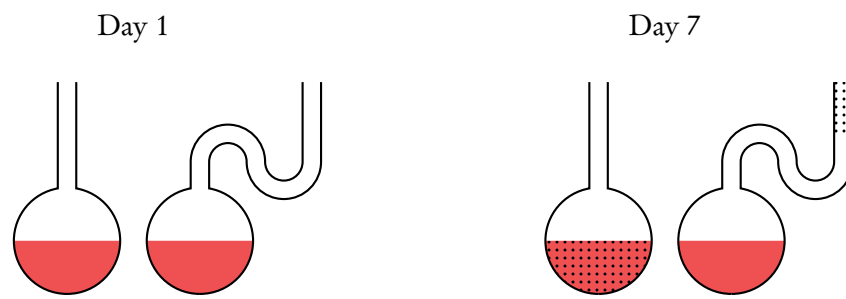
The potato will shrink, its ratio of solute to water will increase and the osmolarities will be balanced.

1.3 Origin of cells

1.3.1 Pasteur's soup

A prior belief was that cells could spontaneously arise from the assembly of inorganic matter. However, Louis Pasteur disputed the belief of spontaneous formation of life in the 19th century.

In his simple experiment, he filled two flasks with nourishing soup, a medium highly nutritious for microorganisms to thrive, and then sterilized them. One flask had a straight open neck, while the other had a curved opened neck. Within a week, the straight-necked soup was spoiled and the curved-necked soup was as good as it was on the first day.



The germs found in the spoiled soup, could be found at the entrance of the curved necked, where they got stuck. Therefore, the mould, fungi and bacteria were able to enter the soup from the environment, but were not able to assemble from thin air in the sealed container.

1.3.2 Formation of organic molecules

In order to form cells, first we have to form (relatively) complex molecules. The Miller-Urey experiment showed that:

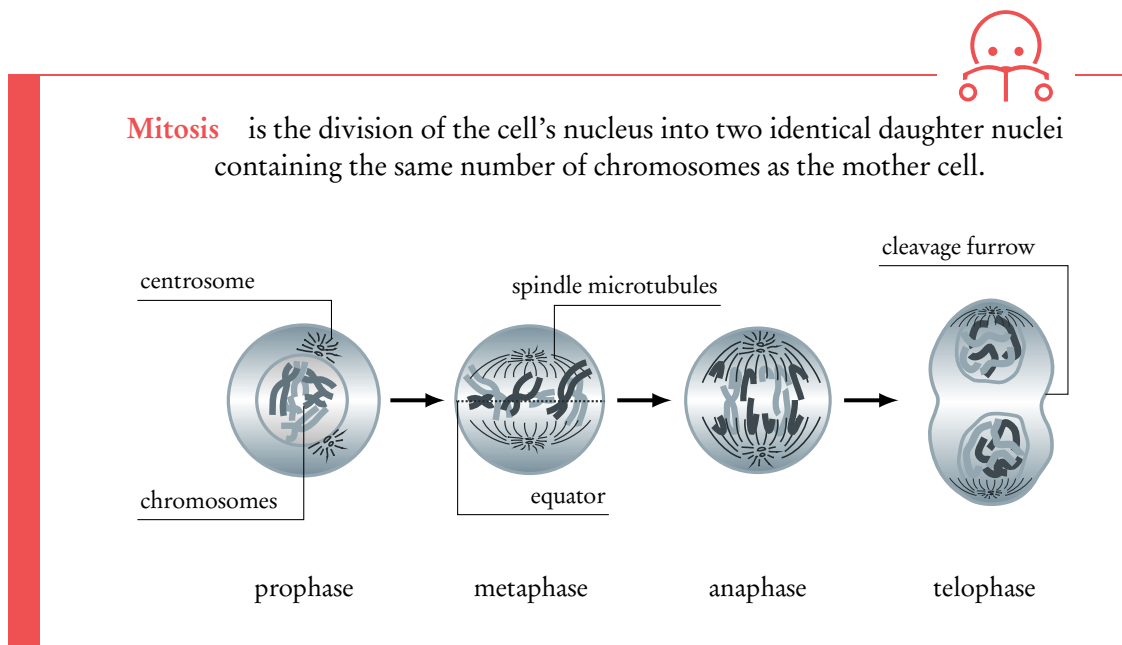
- water vapour, ammonia and methane, all found in the early atmosphere, could have spontaneously assembled into amino acids and carbon compounds, in the presence of electricity (lightning);
- if some of the compounds formed at that time on earth were phospholipids, they would have naturally assembled into bilayers, forming early membranes;
- the formation of nucleic acids such as RNA would have given rise to early enzymatic activities, protein assembly and the first genetic information.

1.3.3 Endosymbiotic theory

Next, this theory assumes that more complex eukaryotic cells have evolved from the prokaryotic cells through a symbiotic process.

- Symbiosis is a mutually favourable coexistence of two organisms.
- The theory suggests that a larger anaerobic prokaryotic cell could have engulfed a smaller aerobic cell, and started coexisting with it.
- The large cell was supplying the smaller one with food, while the smaller cell was converting the food into energy for the larger cell → symbiosis. This would have given rise to mitochondria and chloroplasts.
- Unlike the rest, these organelles are bounded by a double membrane. The two membranes would be the vesicle from endocytosis and the cell membrane of the engulfed cell.
- Mitochondria and chloroplasts contain plasmid-like, circular DNA (characteristic of prokaryotes) that has genes independent from those found in the eukaryotic nucleus. These are thought to be conserved genes from the original engulfed cell.

1.4 Cell division



The function of mitosis is to create two genetically identical daughter cell with the genome of the mother cell. The process involves replication (=duplication) of DNA (all chromosomes). In order for separation of duplicated DNA to work, the DNA (normally

a very long molecule) needs to *supercoil*. Replication is said to be proofread and checked for errors by the cell's machinery.

Remember that mitosis occurs only in eukaryotic cells, while prokaryotic cells divide by binary fission.



Cytokinesis is the division of the cell's cytoplasm and organelles that directly follows mitosis

In plant and animal cells, the process of cytokinesis differs. In plant cell, the kinesis results from the transport of vesicles to the cell equator leading to their eventual fusion and formation of the plasma membrane. The vesicles bring cellulose to form the cell wall around the newly formed plasma membrane. In animal cells, the division of cytoplasm is a result of an invagination of the plasma membrane. Actin and myosin are the contractile fibres that create this invagination called cleavage furrow.

1.4.1 Cell cycle

From its formation, until division, each cell goes through several phases of the life cycle.

- G1 is the phase in which cells spend the majority of their lifespan: this is the period of growth and performance of its daily functions.
- S is the phase that occurs once the cell has decided to undergo mitosis: this is the period of DNA synthesis (replication)
- G2 is the phase where the cell does its last preparations for mitosis: during G2, the cell duplicates its organelles and prepares enzymes and proteins needed for mitosis
- A tip to remember the phases is to know them as G(rowth)1, S(ynthesis of DNA), and G(rowth)2.



Cyclins are proteins that regulate the cell cycle.

The name cyclin should help you remember that the concentrations of these proteins go through cycles or vary throughout the cell cycle in response to internal and external signals. An increase or decrease in the concentration of cyclins will influence the progression of the cell cycle.

- Cyclins comprise the cell cycle checkpoints
- The first cell cycle check point occurs between G1 and S phase
- Another checkpoint occurs during S phase before the beginning of DNA replication
- If the cyclins are not produced or activated, the cell cannot pass a cell cycle checkpoint



Cyclin dependent kinases (CDKs) are enzymes whose activity is dependant on the concentrations of cyclins. It is CDKs which ultimately allow progression through a stage of the cycle via phosphorylation specific molecules.

Despite this tightly regulated cell cycle system, some cells manage to escape the checkpoints and form tumours.



Cancer is the result of uncontrollable cell division and tumours are the aggregates of cancerous cells.

- Mutagens are agents that cause mutations in the DNA.
- Some of these mutations can be missed by proofreading machinery leading to gene mutations
- UV light is a known mutagen that cause high rate of DNA mutations that can often not be repaired
- Oncogenes are genes of each cell that are responsible for normal cell division. They are called oncogenes, because a mutation in these genes can lead to the formation of cancer
- If these genes are mutated, they often lead to cancer
- Proto-oncogenes are oncogenes that in their mutated state become overactivated and promote cell division leading to tumour formation
- Tumour suppressor genes are genes that negatively regulate the cell cycle, so when mutated, they fail to prevent uncontrollable cell divisions
- Metastasis refers to the movement of the primary cancerous cells to a new formation where they continue to form tumours.

1.4.2 Phases of mitosis

Mitosis consists of 4 phases that can be distinguished under the microscope. Due to supercoiling of the DNA, the chromosomes become visible and can be tracked during these phases.

Prophase

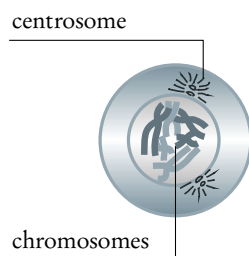


Figure 1.7: Prophase: nuclear envelope is fractured, chromosomes are becoming thick, centrioles are located and the poles of the cell.

- DNA supercoils, chromosomes condense and become visible.
- Nuclear envelope breaks down.
- Spindle microtubules start forming at the poles of the cell.
- The cell contains double the DNA compared to its G1 phase, the same number of chromosomes.
- Begins after S phase has occurred

Metaphase

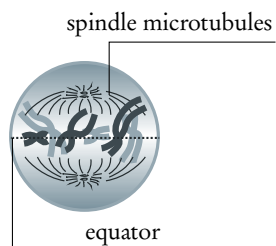


Figure 1.8: Metaphase: chromosomes located at the equator of the cell, each X representing a chromosome, and they're ordered in one single row. Spindle fibres originate at the poles and attach to centres of X-es each pole, with one chromosome having one spindle from each pole.

- Chromosomes align at the equator of the cell.
- Spindle microtubules attach to the centromeres (centres of the chromosome).

Anaphase



Figure 1.9: Anaphase: fibres are shortening towards the poles, and dragging one leg of the X towards the pole. Equal number of chromosome legs are moving to each pole.

- Sister chromatids (legs of each chromosomes, containing identical copies of DNA) are pulled to opposite poles by spindle microtubules.
- Now there is an equal number of chromosomes (DNA molecules) at each pole, but overall, the cell now has double the number of chromosomes compared to prophase.

Telophase

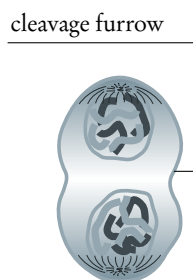


Figure 1.10: Telophase: two nuclei beginning to form at each pole, and the chromosomes uncoiling and becoming longer.

- Chromosomes begin to uncoil as the nuclear envelope reforms around them.
- The cell contains two identical nuclei and awaits the division of cytoplasm and organelles (= Cytokinesis).

When observing a tissue or a group of cells under a microscope, it is easy to calculate the rate of division of the cells/tissue in question. This is done by the following formula:

$$\text{Mitotic index} = \frac{\text{number of cells undergoing mitosis}}{\text{total number of cells}}$$

MOLECULAR BIOLOGY

2.1. Molecules to metabolism 33

- The carbon atom: the core of organic compounds
- Metabolism

2.2. Water 35

- Water: molecular and chemical characteristics – Thermal, cohesive, adhesive, and solvent properties of water
- Hydrophilic vs. hydrophobic substances

2.3. Carbohydrates and lipids 38

- Carbohydrates – Lipids

2.4. Proteins 43




- Amino acids: the building blocks of proteins – The peptide bond: from amino acids to polypeptides – Functions of proteins
- Proteomes: the fingerprints of cells

2.5. Enzymes 46

- Concepts and definitions – Influencing enzyme activity: temperature, pH and substrate concentration

2.6. Structure of DNA and RNA 48

- Nucleotide structure – DNA vs. RNA – The formation of the DNA double helix

	2.7. DNA replication, transcription and translation	51
	– DNA replication – Transcription: from gene to messenger RNA (mRNA) – Translation: making functional proteins from mRNA sequences	
	2.8. Cell respiration	54
	– Cell respiration: substrates and products – Anaerobic cell respiration – Aerobic cell respiration	
	2.9. Photosynthesis	54
	– Photosynthesis – Light spectrum and chlorophyll – Production of oxygen by photolysis – The Calvin cycle: using energy to form carbohydrates and other carbon compounds – Rate-limiting factors of photosynthesis	

2.1 Molecules to metabolism

2.1.1 The carbon atom: the core of organic compounds

The field of molecular biology aims to explain living processes in terms of the chemical substances involved. The most frequently occurring chemical elements in living things are carbon, hydrogen, oxygen and nitrogen. Carbon (C) in particular is a very important element in the study of living things, as:

- all organic compounds contain C (few exceptions like CO₂ and CO)
- C can form four covalent bonds, and thus allows for the formation of a wide variety of stable and complex compounds
- some of these organic compounds essential for life include carbohydrates, proteins, lipids and nucleic acids.

A diagram of these four types of molecules can be found in Figure 2.1.

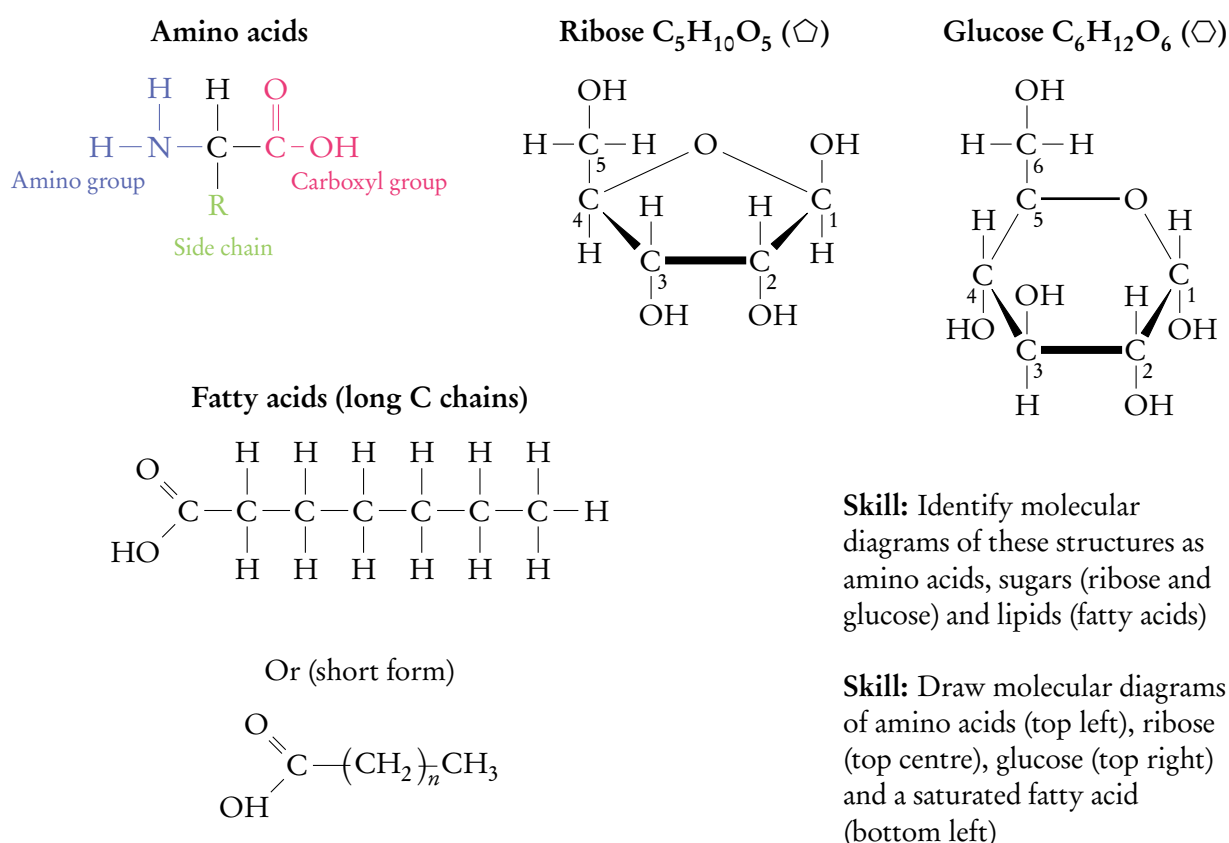


Figure 2.1: Molecular diagrams.

2.1.2 Metabolism

Metabolism is a web of all the enzyme-catalysed reactions in a system (e.g., cell or organism). Metabolic pathways can consist of chains or cycles, and can be anabolic or catabolic.

Anabolism

Synthesis of complex molecules from simpler ones, for example, the formation of macromolecules from monomers by condensation reactions.

Anabolism is associated with condensation reactions, which consist of the removal of a water molecule each time a monomer is added to a polymer chain or another monomer.

E.g., amino acids \rightarrow polypeptide + water.

Catabolism

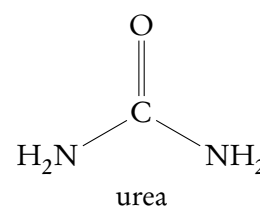
The breakdown of complex molecules into simpler ones, for example, the hydrolysis of macromolecules into monomers.

Catabolism is associated with hydrolysis, which consists of the addition of water molecules to break down a polymer.

E.g., dipeptide + water \rightarrow 2 amino acids.

Example.

Urea: endogenous molecule or artificially produced toxic compound?



In earlier days, organic molecules were believed to be solely synthesized in living organisms. Because urea is an organic compound synthesized in the kidneys as a waste product, it was believed to only be an endogenous molecule. In the early 1800s however, researchers first managed to synthesize artificial urea using silver isocyanate and ammonium chloride. Nowadays it is used as a nitrogen-releasing fertilizer, as well as in the automobile industry and for medical use.

2.2 Water

2.2.1 Water: molecular and chemical characteristics

Water is an essential molecule for life on Earth. It is a polar molecule that consists of 2 hydrogen atoms bound by covalent bonds to an oxygen atom. The principle of covalent bonding consists of sharing of electrons between atoms. It is this bonding gives water its most important characteristic for living organisms: being polar. This polarity arises as water has a slightly positively charged pole where the hydrogen atoms are located and a slightly negatively charged pole where the oxygen atom is located.



Polar molecule is a molecule that has an uneven distribution of charges across the molecule. For example, it is more negative at one end and more positive at another.

Non-polar molecule is a molecule that has an even distribution of charges across the molecule, so no positive or negative poles are formed

Due to the polarity of water molecules, the small negative charge on the oxygen atom has the ability to attract the slightly positively charged hydrogen atoms in nearby hydrogen atoms from other molecules. This attraction leads to the formation of hydrogen bonds between molecules and can explain a number of properties of this molecule, including thermal, cohesive, adhesive and solvent properties.

It is important to remember that in water, covalent bonds are formed between a hydrogen and an oxygen of the same molecule, while hydrogen bonds are found between a hydrogen from one molecule and an oxygen from another

2.2.2 Thermal, cohesive, adhesive, and solvent properties of water

Thermal properties

High specific heat capacity: large amounts of energy are needed to raise the water's temperature. Hydrogen bonds are said to be the strongest of the weak bonds as they restrict movement, meaning it takes a lot of energy to break them down

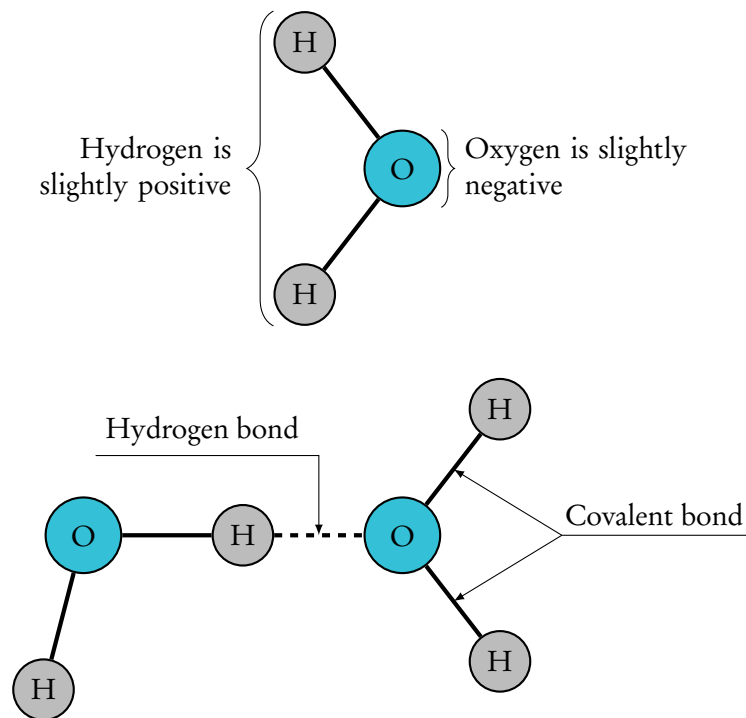


Figure 2.2: Diagrams showing the molecular structure of water (top) and the hydrogen bonds between two water molecules (bottom).

High latent heat of vaporization: hydrogen bonds between water molecules in a liquid form make it very hard for single molecules to escape as vapour. The energy necessary to break these hydrogen bonds and vaporize water is very high compared to other liquids (100 °C). When water vaporizes, a large release of energy occurs, causing a cooling effect on the surface upon which the water used to rest. The concept of sweating as a cooling effect demonstrates this: all the energy used to break hydrogen bonds is released, cooling the skin.

High latent heat of fusion: water at 0 °C must lose a lot of energy before forming ice crystals. Water expands as it freezes and therefore ice can float upon its surface.

Cohesive properties

Water molecules can stick to each other through the formation of hydrogen bonds between the hydrogen of one and the oxygen of another water molecule.

Can explain the formation of water droplets, why some organisms can “walk on water”, etc.

Adhesive properties

Water can adhere to charged surfaces through the formation of hydrogen bonds due to its polarity.

Solvent properties

Water is an excellent solvent for other polar molecules that attract the charged poles of water molecules (e.g., inorganic molecules with positive or negative charges, polar organic molecules, enzymes, etc.). Water can form bonds around other polar compounds, such as NaCl, separating them. Compounds and molecules that dissolve in water are referred to as hydrophilic. Water can also form hydrogen bonds around molecules whose elements are tightly bonded and thus acts as an ideal transport medium for polar molecules (like glucose in blood)

2.2.3 Hydrophilic vs. hydrophobic substances



Hydrophilic (“water-loving”) are all molecules that can readily dissolve in water and can freely associate with it by forming intramolecular bonds. These include polar molecules and ionic compounds

Hydrophilic compounds can readily dissolve in water.

Hydrophobic (“water-hating”) are all molecules that cannot associate with water molecules or easily dissolve in it. These include large and non-polar molecules.

These molecules tend to be insoluble in water.

The hydrophilic and hydrophobic nature of compounds, as well as the solvent property of water, are essential in the transport of molecules in blood for example (which has a high water content). Below is the mode of transport of various important molecules based on their solubility in water:

- Glucose and amino acids are polar, so they can be freely transported and dissolved in blood.
- Cholesterol and fats are non-polar so they are transported in small droplets called lipoproteins, where these non-polar molecules are coated by phospholipids and proteins, which are in turn, polar themselves.
- Oxygen is non-polar, and while some molecules can dissolve in water, they are not sufficient to supply the entire body, therefore, most oxygen is transported in the blood bound to haemoglobin.

A good molecule used to illustrate the importance of the polarity of water and hydrogen bonding for living organisms is methane.

Methane, as opposed to water, has four hydrogens bound to its central atom (in this case a carbon). This causes methane to have an even distribution of charges across the molecule, or a tetrahedral Lewis structure in comparison to water's bent Lewis structure, making it non-polar. This non-polarity gives methane very different properties than those previously discussed for water, and these can demonstrate the vital role of the polarity of water. The latter is shown in Figure 2.3.

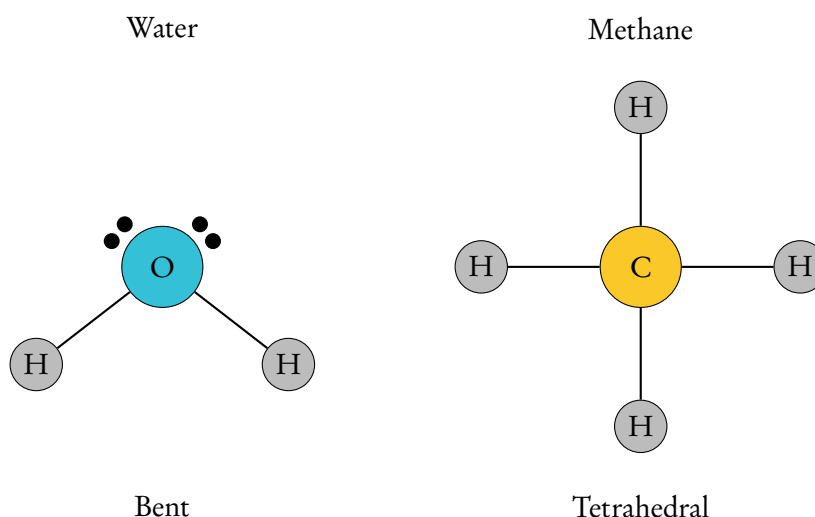


Figure 2.3: Water vs. Methane.

Example.

Comparing thermal properties of water and methane

Property	Methane	Water	Explanation
Melting point	-182°C	0°C	Ice melts at a much higher temperature: hydrogen bonds restrict the movement of water molecules and heat is needed to overcome this.
Specific heat capacity	$2.2\text{ J}/(\text{g }^{\circ}\text{C})$	$4.2\text{ J}/(\text{g }^{\circ}\text{C})$	Water's heat capacity is higher: hydrogen bonds restrict movement so more energy is stored by moving molecules of water than methane.
Latent heat of vaporization	760 J/g	2257 J/g	Water has much higher heat of vaporization: much heat energy is needed to break hydrogen bonds and allow a water molecule to evaporate.
Boiling point	-160°C	100°C	Water's boiling point is much higher: heat energy is needed to break hydrogen bonds and allow water to change from a liquid to a gas

2.3 Carbohydrates and lipids

2.3.1 Carbohydrates

Carbohydrates are organic molecules composed of hydrogen, oxygen, and carbon atoms. Monosaccharides are the monomers of carbohydrates, and are therefore the building blocks of more complex carbohydrates.



Monomer is the building block or basic unit of a class of compounds that can be polymerized to make larger compounds

Dimer is a compound made from the bonding of two monomers

Polymer is two or more repeated monomers from a class of compounds bound together, forming a more complex molecule

The most important carbohydrates from monomers to polymers are shown in the table below:

- monomer = monosaccharide
- dimer = disaccharide
- polymer = polysaccharide

Table 2.1

Monosaccharide	Glucose (G)	Fructose (F)	Galactose (Ga)
Disaccharides	Maltose (G+G)	Sucrose (G+F)	Lactose (G+Ga)
Polysaccharides (all polymers of G)	Cellulose	Glycogen	Starch/Amylose/Amylopectin

2.3.2 Lipids

Lipids are hydrophobic compounds that have important functions in:

- Long term energy storage.
- Heat insulation.
- Buoyancy.
- Shock absorption

The main monomers of lipids are fatty acids: long hydrocarbon chains with a carboxyl group at the end. Fatty acids may be:

Saturated: all the carbon atoms in the fatty acid chain are connected by single covalent bonds, so the number of hydrogen atoms connected to each carbon cannot be increased.

Monounsaturated: there is one double bond between two carbon atoms in the fatty acid chain.

Polyunsaturated: there is more than one double bond between the carbons in the fatty acid chain.

Unsaturated fatty acids can be:

Cis unsaturated: hydrogen atoms are bonded to carbon on the opposite sides of the double bond.

Trans unsaturated: hydrogen atoms are bonded to carbon on the same side of the double bond.

There are three main classes of lipids: phospholipids (important membrane components) steroids (cholesterol and hormones) and triglycerides (long-term energy storage). We will look at the formation of triglycerides, important in energy storage, by means of a condensation reaction.

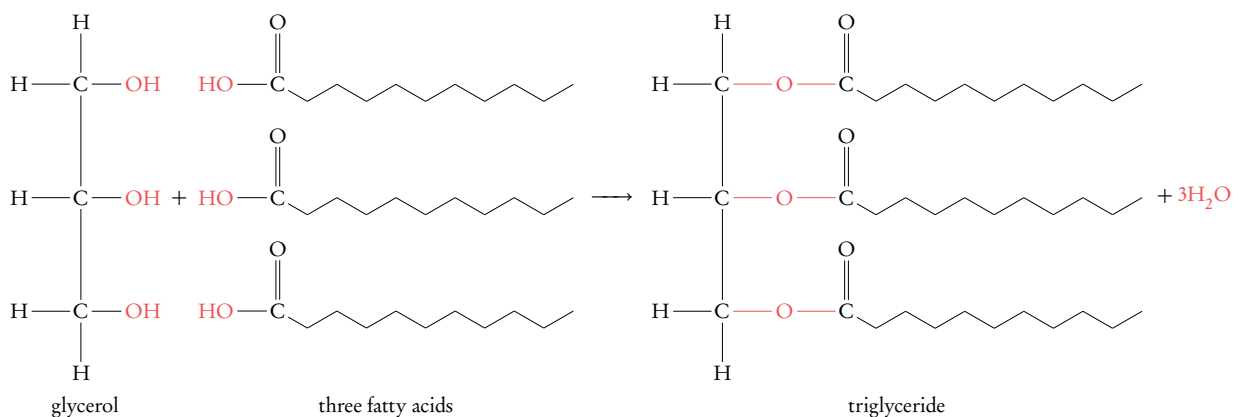


Figure 2.4: Triglyceride formation.

Both lipids and carbohydrates are suitable for energy storage. However, each is more suitable for a specific function:

Carbohydrates

- More easily digested than lipids, good for energy storage that needs to be more rapidly released.
- Soluble in water → easier to transport in blood.

Lipids

- Can store more energy per gram than carbohydrates → better for long term energy storage.
- Not soluble in water, also harder to break down and transport around the body (build-up of high energy content fats).

Example.

Health issues associated to trans- and saturated fatty acids

Trans fats have been banned in several countries in the world, as there is a positive correlation between a diet high in trans fats and coronary heart disease.

Saturated fats have also been shown to have a positive correlation (albeit, weaker than trans fats) with the incidence of coronary heart disease.

However, many of the tested populations do not fit these findings, so evidence must be carefully evaluated before banning products and establishing anti trans or saturated fat campaigns.

Determining the body mass index (BMI)

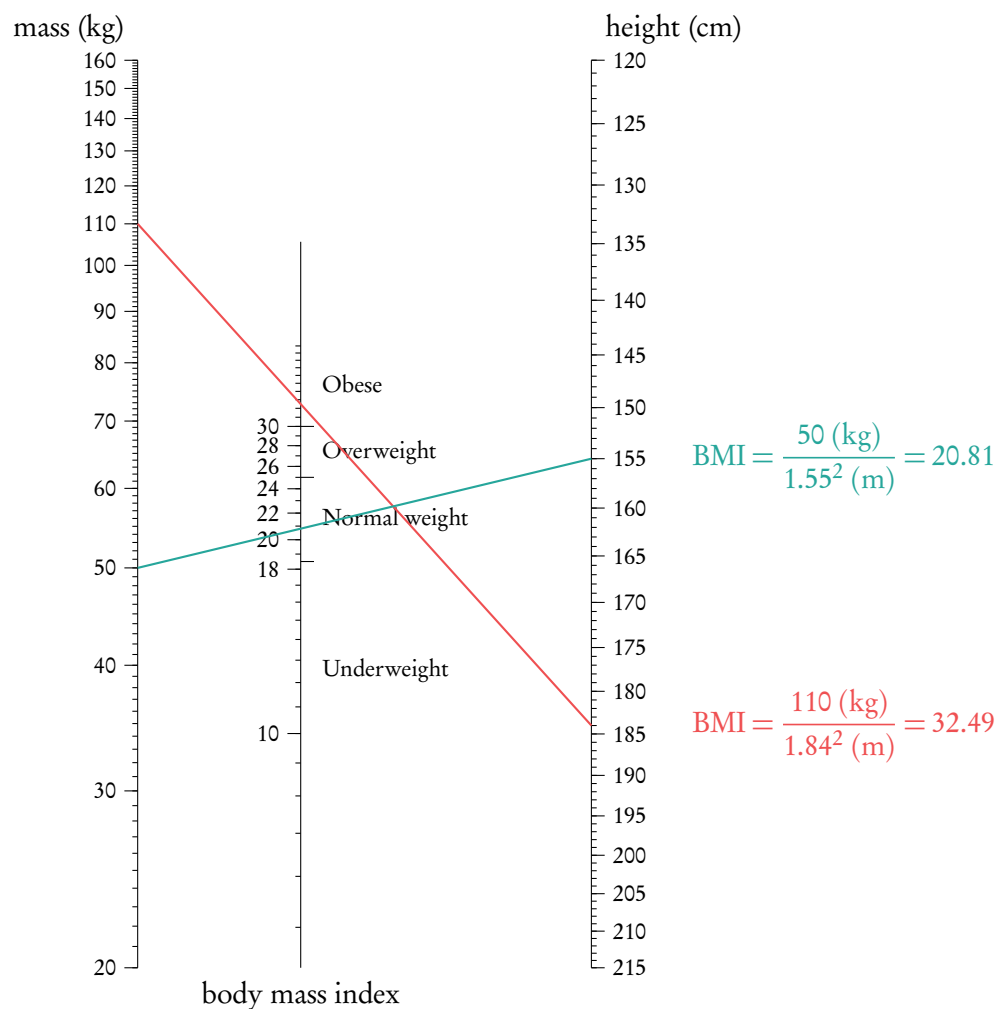
Because of the natural variation in size between adults, weighing someone does not provide a clear indicator of body mass. The body mass index is a screening tool to identify possible weight problems, and it can be calculated using the following formula:

$$\text{BMI} = \frac{\text{mass in kg}}{(\text{height in meters})^2}$$

where:

Body mass index	Conclusion
Below 18.5	Underweight
18.5 to 24.9	Normal weight
25.0 to 29.9	Overweight
30.0 or more	Obese

A nomogram can also be used to calculate BMI, by drawing a line that connects the height and weight lines, the BMI measure is indicated by the scale in the middle.



2.4 Proteins

2.4.1 Amino acids: the building blocks of proteins

Amino acids, containing a carboxyl, an ammine and an R group, are the monomers of proteins that when linked together by peptide bonds form complex proteins. Proteins are important organic molecules that carry out major functions in cells and in the extracellular space.

Although there are only 20 different amino acids, millions of proteins exist as these monomers can be linked in any given sequence

This means that if you have a protein made of n amino acids, there are 20^n different proteins that may be made.

The specific sequence of each protein is coded for in the genetic material of the organism. As we will see later in the chapter, DNA is transcribed into mRNA and later translated by ribosomes into polypeptide chains.



The Central Dogma of molecular biology states that there is a sequential transfer of information where DNA is transcribed into RNA, which in turn is translated into proteins.

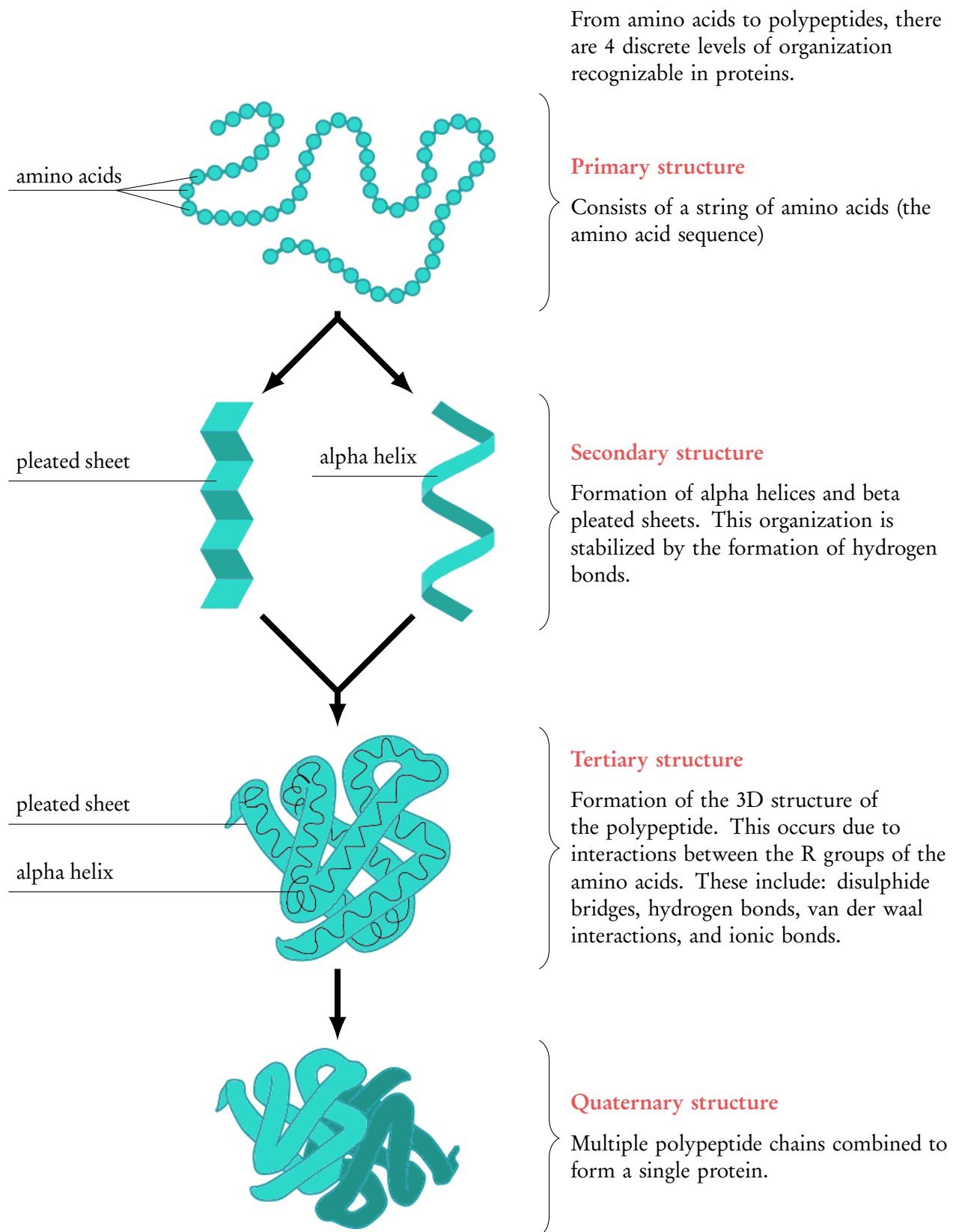


Figure 2.5: Protein organisation.

2.4.2 The peptide bond: from amino acids to polypeptides

Figure 2.6 is a diagram showing the formation of a dipeptide (a 2-amino acid molecule) via a condensation reaction, and the breakdown of a dipeptide into two amino acids. The condensation reaction creates a covalent, peptide bond between the carboxyl group of one amino acid and the amino group of the other, and results in the release of a water molecule. In the hydrolysis reaction (Figure 2.7), water is added in order to break the peptide bond.

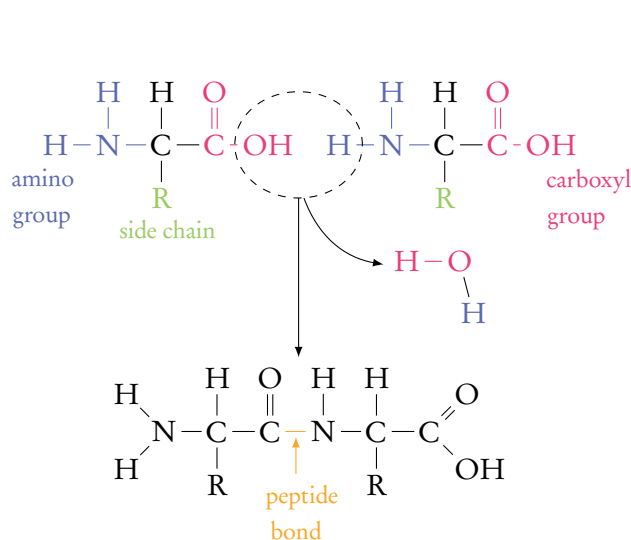


Figure 2.6: Condensation.

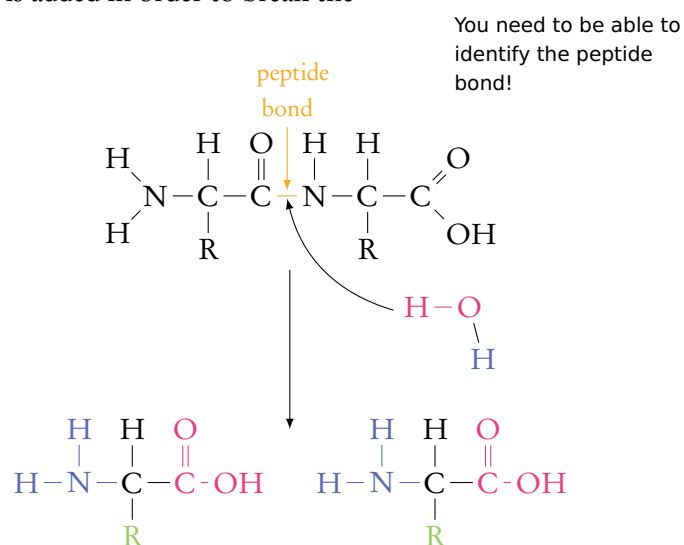


Figure 2.7: Hydrolysis.

2.4.3 Functions of proteins

Table 2.2 depicts some of the major functions proteins carry out in an organism, as well as specific examples for each function.

Table 2.2: Protein functions.

Function	Example	Details	Shape
Structural	Collagen	Strengthen bone, tendon and skin	Fibrous
Transport	Hemoglobin	Bind oxygen in the lungs and transports to other tissues	Globular
Movement	Actin	Involved in the contraction of muscles	Fibrous
Defence	Immunoglobulins	Acts as antibody	Globular

2.4.4 Proteomes: the fingerprints of cells

Proteome is the entire set of proteins expressed by a genome, cell, tissue, or organism at a given time. While the genetic make up of an organism is the same in all cells, each tissue or individual cell shows variable gene expression and thus different proteins are created. The proteome of individuals within the same species is quite similar (as the genetic make up is also similar), however, each individual has a unique proteome (like a fingerprint, which can be similar but never identical to other individuals).

2.5 Enzymes

2.5.1 Concepts and definitions



Enzymes are globular proteins that function as biological catalysts that speed up chemical reactions in biological processes.

- Substrates are substances acted upon by enzymes.
- Active site is the region on the enzyme to which substrates bind and where catalysis occurs.
- The activity of enzymes relies on the concepts of molecular motion and collision, in other words, substrates and enzymes must “collide” with one another due to their individual motion (kinetic energy). The more collisions between enzyme and substrate, the faster the reaction occurs.
- Enzymes speed up reactions without getting consumed by the process, meaning they can speed up many reactions.

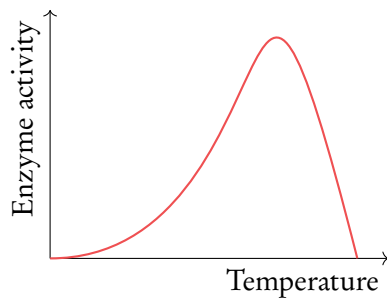
There are two main models that aim to explain the mechanism of action of enzymes:

The Lock-and-key model: the substrate and enzyme have shapes that make them fit perfectly with each other. Thus, each enzyme catalyses a specific reaction

The Induced-fit model: as substrate and enzyme approach each other, their interactions make them shift physical conformation so that they fit perfectly with one another.

2.5.2 Influencing enzyme activity: temperature, pH and substrate concentration

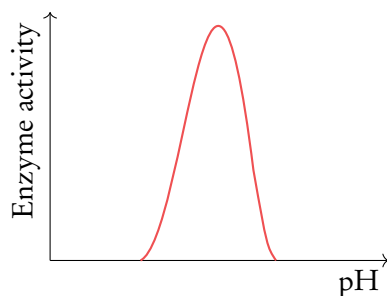
Temperature



Enzyme activity increases as temperature increases, often doubling with every 10 °C rise. This is because collisions between substrate and active site happen more frequently at higher temperatures due to faster molecular motion.

Enzymes are proteins, therefore at high temperatures they are denatured and stop working. This is because heat causes vibrations inside enzymes which break bonds needed to maintain the structure of the enzyme.

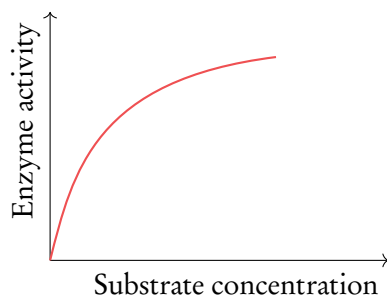
pH



Enzyme activity is reduced as pH increases above the optimum because the conformation of the enzyme is altered more and more. Above a certain pH the alkalinity denatures the enzyme and it does not catalyze the reaction at all.

Enzyme activity is reduced as pH increases above the optimum because the conformation of the enzyme is altered more and more. Above a certain pH the alkalinity denatures the enzyme and it does not catalyze the reaction at all.

Substrate concentration



At low substrate concentrations, enzyme activity increases steeply as substrate concentration increases. This is because random collisions between substrate and active site happen more frequently with higher substrate concentrations.

At high substrate concentrations most of the active sites are occupied, so raising the substrate concentration has little effect on enzyme activity. A plateau is reached when enzymes are working at full capacity at their maximum rate

Example.

The use of lactase in the production of lactose-free milk

Many enzymes are used in industrial processes (for instance, in the food industry). Enzymes are often immobilized on a surface and employed in large concentrations to catalyse a wide range of biochemical reactions. A common example is the use of enzyme lactase in the production of lactose-free milk.

Lactose is the disaccharide in milk that many people are intolerant to as they do not produce the enzyme lactase to break it down. Often times milk and other milk products are treated with immobilized lactase, and lactose is broken down prior to consumption. The resulting monosaccharides are easier to digest by lactose-intolerant people, and result in a sweeter flavour (less artificial additives needed). The use of the enzyme also speeds up the production of fermented products like yogurt and cheese.

Immobilized lactase can be used in much larger concentrations and can resist larger changes in pH and temperature compared to endogenous lactase.

The immobilized enzymes can also be reused, and the products are not contaminated with enzymes (easier to introduce and remove from the sites of reaction).

2.6 Structure of DNA and RNA

2.6.1 Nucleotide structure

Nucleic acids are the biomolecules responsible for information storage, essential to all forms of life. The two major types of nucleic acids DNA (deoxyribonucleic acid) and RNA (ribonucleic acid) are essential compounds involved in gene expression in cells. RNA and DNA polymers consist of repeated units of nucleotides, which are in turn made of a 5 carbon sugar linked to a phosphate group at carbon 5, and to one of five nitrogenous bases (adenine, guanine, thymine, uracil and cytosine) at carbon 1. The overall nucleotide structure is shown in Figure 2.8.

Skill: Draw a simple diagram of the structure of single nucleotides (you may use simple circles for the phosphate, pentagons for the sugar [deoxyribose or ribose] and rectangles for the base).

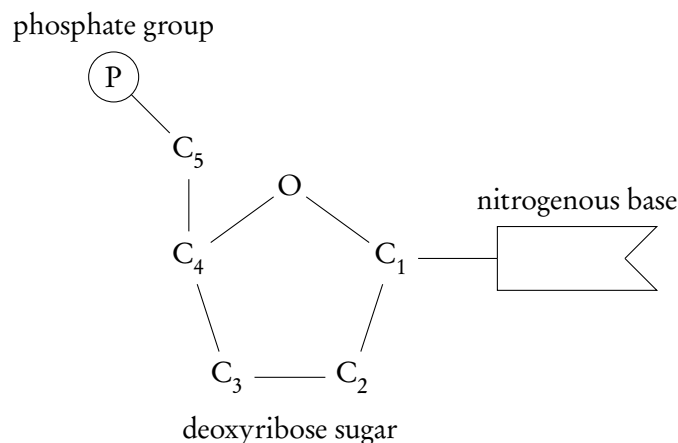


Figure 2.8: Nucleotide structure.

The nucleotide can have either a ribose (RNA) or a deoxyribose (DNA) pentose sugar. These differ in the presence or absence of an oxygen molecule. This oxygen molecule makes ribose a less stable molecule than deoxyribose, due to the fact that oxygen has high electronegativity, meaning that it really wants to more electrons. This instability causes RNA to be single stranded while DNA can be double stranded.

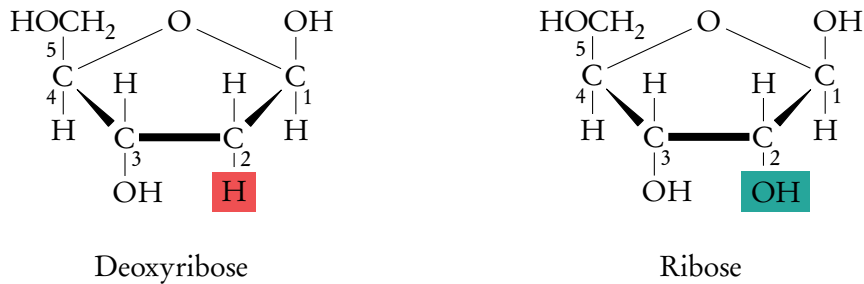


Figure 2.9: Deoxyribose and ribose.

2.6.2 DNA vs. RNA

Both types of nucleic acids share structural similarities, but also significant differences:

RNA

- Contains a 5-carbon sugar
- Sugar is called ribose
- Single-stranded molecule
- Contains bases adenine (A), uracil (U), cytosine (C), and guanine (G)

DNA

- Contains a 5-carbon sugar
- Sugar is called deoxyribose
- Double-stranded molecule
- Contains bases adenine (A), thymine (T), cytosine (C), and guanine (G)

2.6.3 The formation of the DNA double helix

DNA is composed of a double stranded helix of DNA nucleotides. Each strand of DNA is held together by covalent bonds that form between the phosphate group of one nucleotide to carbon 3 of the neighbouring nucleotide. This forms a single-stranded backbone. The DNA double strand is then achieved by the formation of hydrogen bonds between the nitrogenous bases of two nucleotide strands. Base pairing in DNA is complementary, meaning that one base can only bind to a specific complementary base:

- Adenine (A) binds to thymine (T) → 2 hydrogen bonds.
- Cytosine (C) binds to guanine (G) → 3 hydrogen bonds.

The two DNA strands are antiparallel, in other words, they run in opposite directions (where one strand has a 5' end, the complementary strand has a 3' end).

The diagram shows the structure of the DNA double helix, showing two joined antiparallel DNA strands bound together by complementary base pairing of adenine with thymine (2 H-bonds); and cytosine and guanine (3 H-bonds).

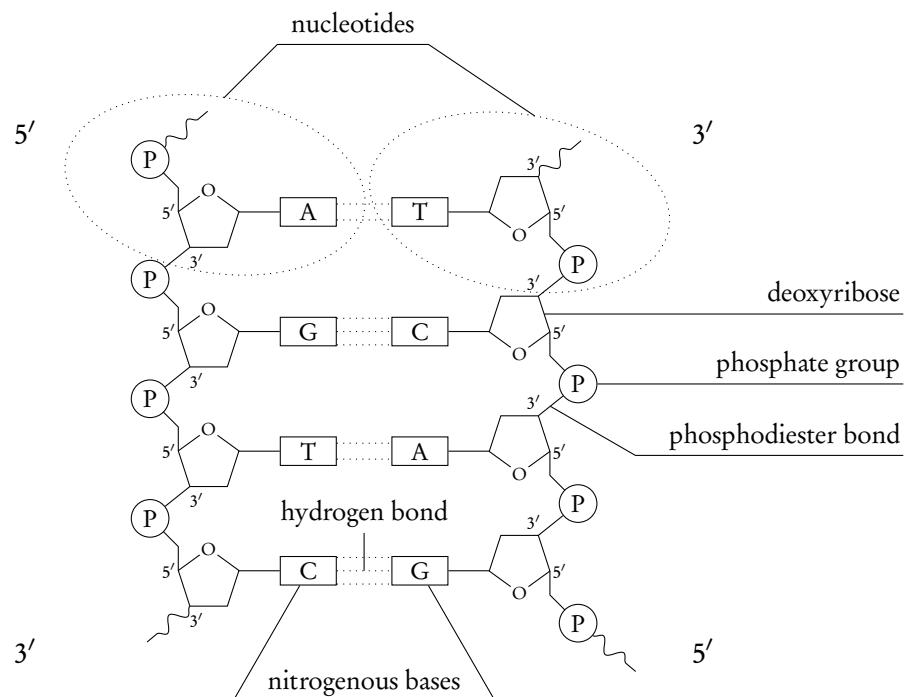


Figure 2.10: DNA structure.

2.7 DNA replication, transcription and translation

2.7.1 DNA replication

During the DNA replication process, one double stranded DNA molecule gives rise to two daughter DNA molecules. This process is said to be semi-conservative, meaning that each new DNA double helix contains one newly synthesized daughter strand and one strand from the original parent DNA strand, which serves as a template to ensure that both new strands are identical. Essentially, part of the original DNA is conserved at each replication step.

Below is a brief description of the process of DNA replication:

- Takes place during the synthesis (S) phase of the cell cycle.
- Helicase unwinds the double helix and separates the two DNA strands by breaking hydrogen bonds.
- The two parent strands that emerge from this process serve as templates for the new daughter strands to be synthesized.
- Enzyme DNA polymerase can then link free nucleotides to the template strands by complementary base pairing. Note that DNA polymerase can only add nucleotides at the 3' end of a growing strand.
- Two identical daughter DNA strands are created, resulting in two semi-conservative double stranded DNA helices.

Example.

Taq DNA polymerase: production of multiple copies of DNA by polymerase chain reaction (PCR)

This technique has been one of the greatest biotechnological developments in DNA research. It allows scientists to amplify desired regions of DNA in very little time. PCR consists of the following steps:

1. Isolate the desired region of DNA (using restriction enzymes).
2. Introduce it in a mixture containing free nucleotides, primers and Taq DNA polymerase.
3. The mixture is heated up to 90 °C to separate the DNA strands of the original template.
4. Temperature is then reduced to 55 °C to allow for primer annealing to the now separated strands.
5. Taq polymerase (isolated from thermophiles, organisms that can survive at very high temperatures) works optimally at 72 °C, so the mix is heated to this temperature to enhance the formation of new double-stranded copies of the original DNA.
6. Process is repeated several times until the DNA is amplified.

2.7.2 Transcription: from gene to messenger RNA (mRNA)

Transcription is the synthesis of mRNA copied from the DNA base sequences present in an organism's chromosomes. The sections of DNA that code for polypeptides are called genes, but in order for these polypeptides to be expressed, machinery located outside the nucleus is needed. Thus, a messenger RNA (mRNA) molecule carries the “message” from the DNA to the cytoplasm. Below is the explanation of the process of transcription:

Skill: While you must be able to determine the mRNA sequence that will result from a given DNA sequence, it is essential that you can also go backwards, and deduce the original DNA sequence from a given mRNA sequence. Try it with the following mRNA sequence!

Tip: first deduce the complementary base sequence to the given mRNA to get one of the DNA strands, the second DNA strand, as you will realize, is identical to the mRNA sequence, with thymine replacing the uracil bases in the sequence.

- RNA polymerase unwinds the area of the DNA to be transcribed.
- RNA polymerase catalyses the addition of free RNA nucleotides using one of the newly separated DNA strands as a template for complementary base pairing (this creates a copy of the complementary DNA strand containing the gene of interest).
- In this process, thymine is replaced by nitrogenous base uracil (only present in RNA nucleotides).
- Translation occurs in a 5' to 3' direction.
- Once the whole gene has been transcribed, the resulting single-stranded mRNA molecule peels off and moves out of the nucleus to be translated into a polypeptide.

A U C G A A C G U U G G G C C C G A

2.7.3 Translation: making functional proteins from mRNA sequences

Once the DNA “message” has entered the cytoplasm in the form of mRNA, translation takes place, where polypeptides are synthesized by ribosomes. When studying this process, researchers found that the genetic code is written in a language of codons: which consist of three consecutive bases (triplet), where each codon codes for a specific amino acid (see the table below to see which codons code for what amino acids). Codons are located on the mRNA sequence, while anticodons (complementary to codons) are found on tRNA molecules, another type of RNA that carries the appropriate amino acid to the ribosome where translation occurs. Below is a description of this process:

- The mRNA strand created during the process of transcription binds to a ribosome.
- The ribosome begins to slide over the mRNA until it reaches a starting codon, where a tRNA with a complementary anticodon can bind, bringing the first amino acid of the polypeptide to be made.
- A second tRNA molecule with the appropriate anticodon binds to the second codon.
- The ribosome catalyses the formation of a peptide bond between the two amino acids, creating a dipeptide carried by the second tRNA.
- The ribosome slides over the mRNA molecule, leading to the release of the first tRNAs (the one that is no longer carrying an amino acid) and the binding of a new tRNA to the following codon.

- The amino acid chain keeps growing as this process is repeated until a stop codon is reached, at which point the polypeptide breaks away from the tRNA and can fold and be modified to become a functional protein.

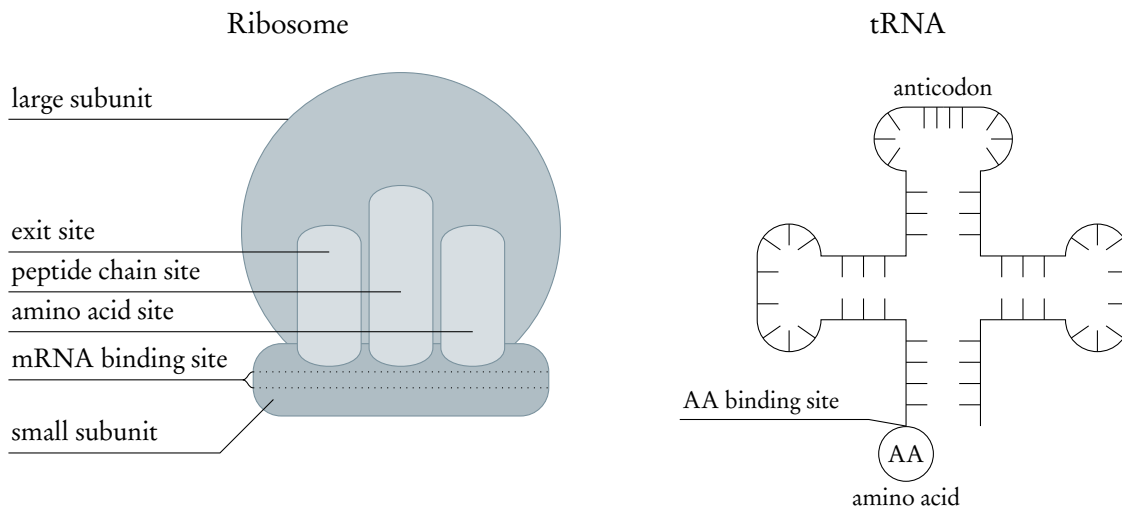


Figure 2.11: Ribosome structure and tRNA.

Example.

Universality of the genetic code: producing human insulin in bacteria

The genetic code has been shown to be universal. That is, in all organisms, the codon code is the same (one codon codes for the same amino acid in any organism). This is very advantageous, as researchers have been able to synthesize important proteins at higher rates by introducing a human DNA sequence for instance in a smaller organism like *E. coli*, resulting in faster synthesis of the desired protein.

Below is a table showing this universal codon code:

Insulin is a great example of this. Researchers isolated the insulin gene from humans and introduced it into *E. coli* (a bacterium that rapidly replicates and can yield large amounts of protein in very short time periods). *E. coli* can then transcribe and translate the insulin gene using its innate machinery. Researchers can then isolate and purify this very important enzyme and use it in for example, the treatment of diabetic patients.

ACTACGTACCTGGGACTAGACT

Skill: Use a table of the genetic code to deduce which codon(s) corresponds to which amino acid. For example, try coding the following DNA sequence (only one strand is given) into its transcribed mRNA sequence and this sequence into separate amino acids (remember to first find the start codon, and to correctly identify the stop codon, if present).

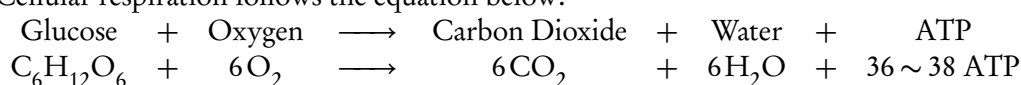
2.8 Cell respiration

2.8.1 Cell respiration: substrates and products



Cellular respiration is the controlled release of energy, in the form of ATP, from organic compounds in cells

Cellular respiration follows the equation below:



Cell respiration can follow an aerobic (in the presence of oxygen) and an anaerobic pathway (no oxygen). The latter creates a much smaller yield of ATP.

2.8.2 Anaerobic cell respiration

When no oxygen is available to the cells, the following process occurs:

- Glycolysis occurs in the cell's cytoplasm, where a glucose molecule is broken down into two smaller 3-carbon molecules called pyruvate.
- This process leads to a small yield of ATP (2 molecules per reaction) and other products that can later be used in aerobic cell respiration.
- In yeast cells, pyruvate is converted into ethanol and carbon dioxide (there is no further yield of ATP and the products are released as waste). This process is known as fermentation.
- In mammalian cells, pyruvate molecules are converted into lactate molecules (also known as lactic acid), with no further yield of ATP. Lactate accumulates and can lead to changes in pH (lactic acidosis), which can be dangerous in the long term

2.8.3 Aerobic cell respiration

When oxygen is present, pyruvate can be further broken down in the cytoplasm and enter the mitochondria in the form of acetyl-CoA (a 2-carbon molecule).

- Acetyl-CoA enters the Krebs cycle, where a series of redox reactions lead to the release of carbon dioxide and the formation of intermediate molecules.
- These molecules are used in the electron transport chain (at the mitochondrial membrane), resulting in a large yield of ATP (34-36 ATPs) and the release of water as a by-product

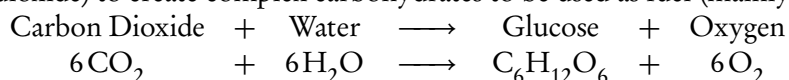
2.9 Photosynthesis

2.9.1 Photosynthesis



Photosynthesis is the process in which plants produce their own organic substances to be used as nutrients.

The process uses energy from the Sun and simple organic compounds (water and carbon dioxide) to create complex carbohydrates to be used as fuel (mainly glucose) and oxygen:



2.9.2 Light spectrum and chlorophyll

Sunlight is made up of a range of wavelengths including colours red, green and blue within the visible light spectrum. The smaller the wavelength the more energy is reflected (blue wavelength), and the larger the wavelength the less energy reflected (red). Green colour is reflected from medium wavelengths. To absorb and reflect these light waves, specific pigments in plants are needed. The main photosynthetic pigment is chlorophyll; it absorbs red and blue light very well, and reflects mostly green light (thus giving plants their green colour). Chlorophyll is located in clusters inside chloroplasts. Figure 2.12 shows the absorption spectrum of chlorophyll, showing peaks at the wavelengths easily absorbed by the pigment (blue and red) and a trough on green, the least absorbed wavelength. By looking at the action spectrum (Figure 2.9.2, wavelengths of light most used during the photosynthesis reactions) it is clear why chlorophyll is the main pigment in this process: the wavelengths readily absorbed by chlorophyll are majorly used in photosynthesis.

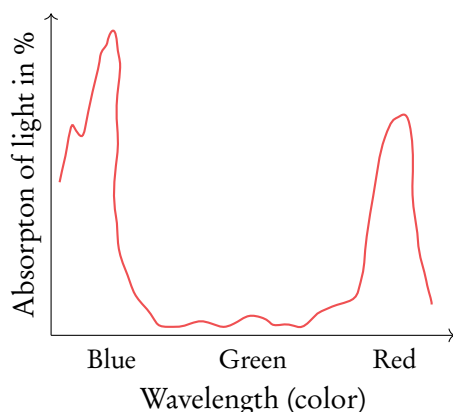


Figure 2.12: Absorption.

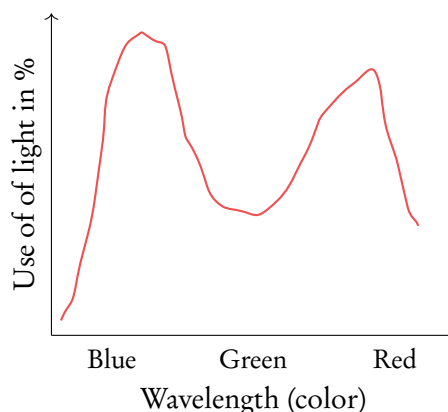


Figure 2.13: Action.

Skill: Draw the absorption spectrum of chlorophyll and the action spectrum for photosynthesis.

2.9.3 Production of oxygen by photolysis

Photosynthesis consists of light-dependent and light-independent reactions. The light dependent reactions result in the yield of ATP, oxygen and hydrogen.

Hydrogen and electrons are then involved in the electron transport chain which results in a yield of ATP and intermediate molecules for the light-independent reactions

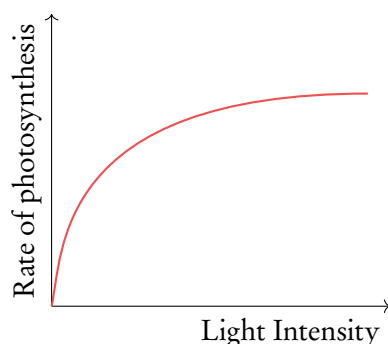
2.9.4 The Calvin cycle: using energy to form carbohydrates and other carbon compounds

The light-independent reactions lead to the formation of complex carbohydrates.

Also known as the Calvin cycle, where ATP and carbon dioxide are used to convert inorganic compounds into organic compounds. This is achieved by carbon fixation, which requires energy from ATP.

2.9.5 Rate-limiting factors of photosynthesis

Light Intensity



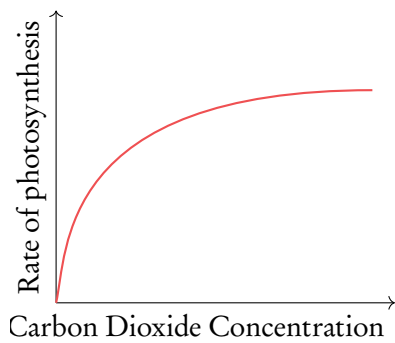
At low light intensities, rate of photosynthesis is limited.

Photolysis, which requires the absorption of light waves slows down, and thus, so does oxygen and ATP production.

Indirectly limits the light-independent reactions, as ATP is necessary for carbon fixation to occur.

The graph levels off once all the enzymes and reactions are occurring at the highest speed possible.

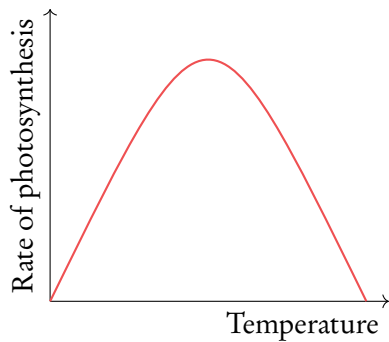
Carbon Dioxide Concentration



Rate-limiting step in the Calvin cycle → carbon cannot be fixed to inorganic compounds and thus glucose production slows down.

Increasing CO_2 concentration increases the rate of photosynthesis, until the photosynthetic enzymes involved in the cycle (e.g., rubisco) reach their saturation point and can no longer increase reaction rates.

Temperature



At low temperatures, the enzymes involved in photosynthetic reactions work very slowly.

Rate of reaction increases steadily as temperature increases, until reaching an optimum point when all enzymes are working at a high rate.

When the temperature surpasses this optimal point, enzymes can be denatured, once again decreasing the photosynthetic rate.

Skill: Design experiments to investigate the effect of these factors on photosynthetic rates.

3.1. Genes and chromosomes **60**

- Genes – Sickle cell anaemia as an example of a gene mutation
- Genome and Human Genome Project
- Complexity of organisms and their genomes
- Prokaryotic and eukaryotic chromosomes – Karyograms and karyotypes

3.2. Meiosis **66**

- Meiosis I – Meiosis II – Meiosis and variation
- Failures of meiosis

3.3. Inheritance **73**

- Mendel's law of inheritance – Punnett grids
- Co-dominant alleles – Sex linkage – Dominant and recessive genetic disorders
- Pedigree charts – Dihybrid crosses
- Linked genes – Crossing over and linked genes
- Polygenic inheritance

3.4. Genetic modifications and biotechnology **84**

- PCR and gel electrophoresis – Genetic modifications
- Benefits and risks of genetic modifications – Clones and cloning

3.5. Gene pools and speciation **90**

- Directional, stabilizing and disruptive selection

3.1 Genes and chromosomes

3.1.1 Genes



Gene a DNA sequence that defines a certain heritable characteristic.

One DNA molecule contains many genes, but not all of these genes are “switched on”, meaning expressed.

Chromosome a DNA molecule that carries genes.

Within a species, all chromosomes are made of the same DNA molecule, with certain variations of alleles.

The expression of genes on this DNA molecule is what differs one chromosome from another.

The number of chromosomes is defined per species. In eukaryotic organisms, chromosomes come in pairs with two chromosomes of the same pair carrying the same genes: a human, for example, has 46 chromosomes, meaning 23 pairs.



Allele a variation of a certain gene, differing from the other allele of the same gene by a few bases. Different alleles code for different variations of the trait coded for by that gene.

Alleles are a result of mutations of the gene sequence. Most genes come in two or more allelic forms. Since chromosomes come in pairs, an organism can have two or more possible alleles of a gene.

If an organism has two of the same alleles, it is called *homozygous* for the trait.

If an organism has two different alleles, it is said to be *heterozygous* for the trait.

Alleles can be dominant and recessive, where **dominant alleles** are always expressed if present, while **recessive alleles** are only expressed if present homozygously.

3.1.2 Sickle cell anaemia as an example of a gene mutation

Sickle cell anaemia is a heritable disease caused by a mutation of a gene coding for the haemoglobin molecule.

A mutation is a change in the base sequence of a DNA molecule.

In sickle cell anaemia, a base substitution mutation causes an adenine base in the GAG triplet to be substituted by thymine (GTG).

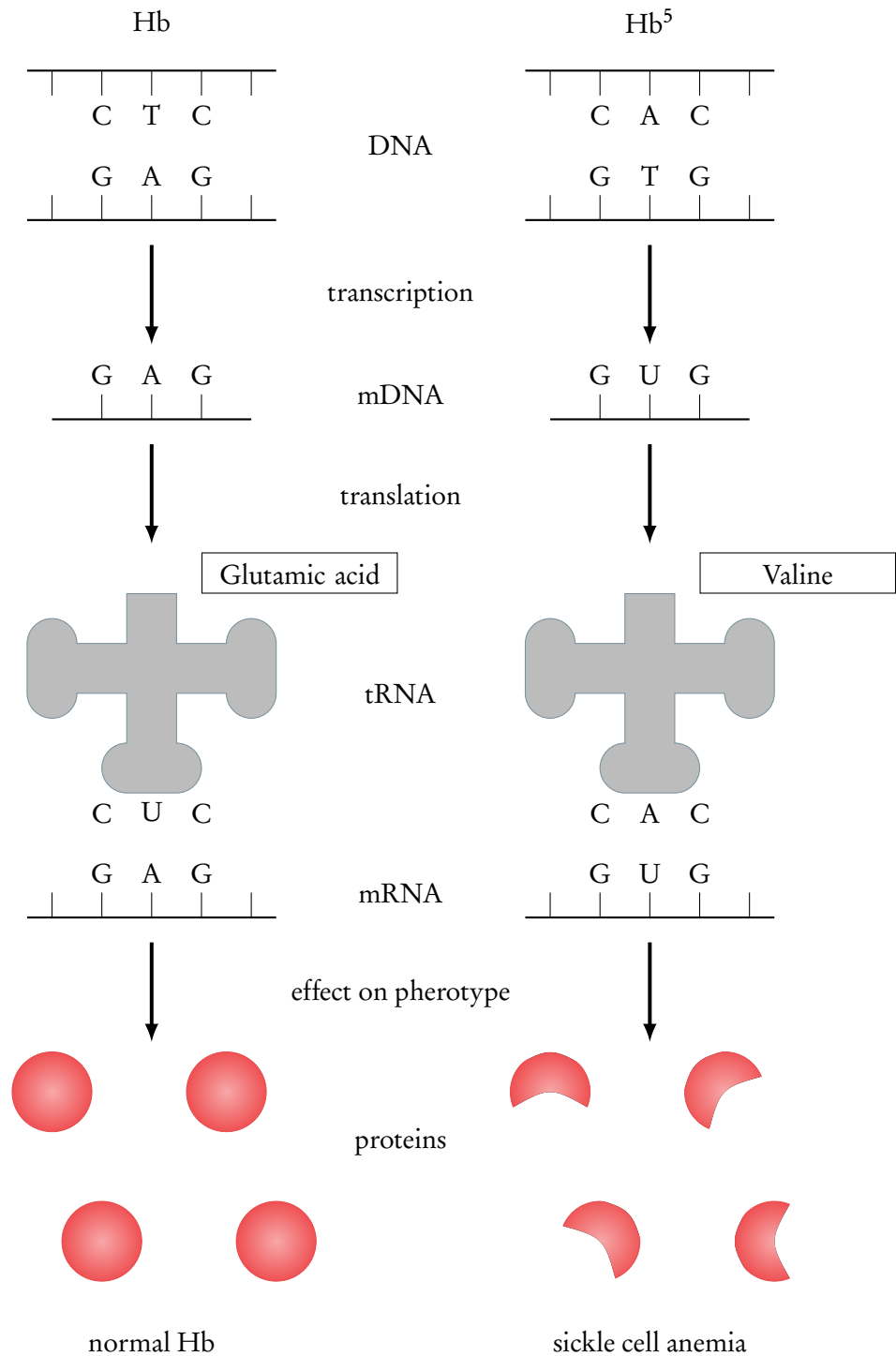
Recall that a triplet of nucleotides codes for a specific amino acid within a protein.

In this case, GAG codes for glutamic acid which becomes substituted by valine (GTG) in sickle cell anaemia.

Recall that protein folding ultimately depends on the amino acid sequence. The difference in this amino acid (from glutamic acid to valine) changes the shape of the haemoglobin protein, leading to a less functional molecule.

Individuals with sickle cell anaemia have moon shaped red blood cells with lower oxygen-carrying capacity compared to normal red blood cells.

On the other hand, malaria parasite is less likely to infect the sickle cells compared to the healthy ones, so the people with this condition are resistant to malaria.



3.1.3 Genome and Human Genome Project



Genome refers to the entire genetic information of an organism.

The genome not only includes genes, but all the genetic information found in the cells of an organism: introns and exons

Animal cell genome includes chromosomal DNA as well as DNA in the mitochondria, just as a plant genome includes all chromosomes and chloroplast DNA.

Bacterial cell genome includes their chromosomal DNA and their plasmid DNA



Human Genome Project refers to the sequencing of the entire human genome.

The HGP resulted in the mapping of the location, size, and number of genes in humans (19 to 20 thousand of them). The project also concluded that humans share most of their genetic sequence, with the exception of short nucleotide polymorphisms (SNPs).

The rest of the DNA sequence is the non-coding region which can control expression of the coding regions: the introns.

3.1.4 Complexity of organisms and their genomes

The size of the genome of a species does not relate to the complexity of the organism.

Similarly, the number of genes in an organism does not correlate with the complexity of that organism

Table 3.1

Organism	Genome size (number of bases)	Number of genes	Number of chromosomes
E. coli	4.6 million	2,300	...
Maize (corn)	2.3 billion	32,000	10
Mouse	2.8 billion	23,000	40
Human	3.0 billion	20,000	46

3.1.5 Prokaryotic and eukaryotic chromosomes

Cairn's method of measuring DNA length

Cairn's method involves adding a radioactively labelled nucleotide (thymidine) to cells undergoing replication.

Recall that during replication, each parent strand gains a new daughter strand which assembles from the available nucleotides through complementary base pairing.

In this way, the two new daughter strands incorporate the radioactive nucleotides (thymidine, which also discriminates between DNA and RNA replication) and can be visualized after light exposure.

This method was used to show the nature of replication in bacterial, and subsequently in animal cells.

Table 3.2: Comparison of prokaryotic and eukaryotic DNA.

	DNA	Chromosomes	Plasmid
Prokaryotic	Circular "Naked"	1 circular	Present
Eukaryotic	Linear Associated with proteins	Many linear	Not present

Prokaryotic DNA

- Bacteria have one circular DNA molecule that is not associated with proteins.
- Extra genetic information is stored on plasmids, and can easily be shared between bacteria.
- Antibiotic resistance genes are often found on plasmid DNA

Eukaryotic DNA

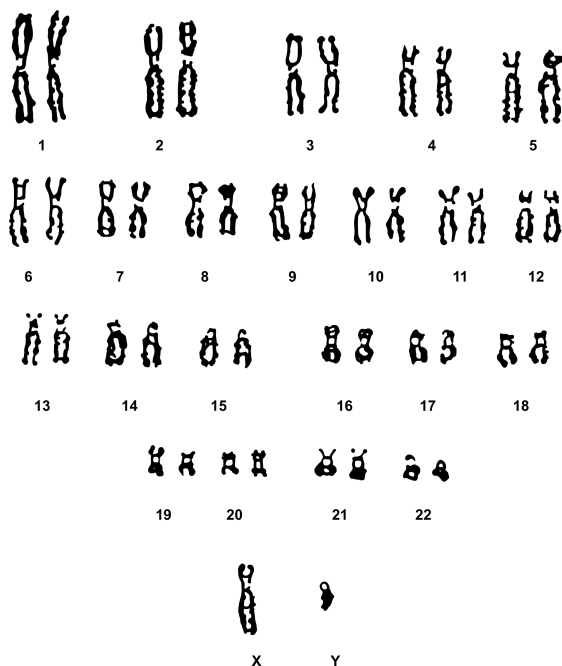
- DNA is associated with proteins called histones.
- Histones are used to wrap DNA around them in order to protect it against damage as well as to control expression of certain genes.
- Recall that eukaryotic chromosomes come in pairs.
- A pair of identical chromosomes is called a homologous pair and these chromosomes carry the same genes (with possibly different alleles).
- A complete set of chromosomes (in a human 46) is called a diploid number of chromosomes.
- Sex cells contain half the number of chromosomes (one from each pair), which is said to be haploid, in order to conserve the species' number of chromosomes after fertilization (joining of two sex cells).

3.1.6 Karyograms and karyotypes



Karyogram is an image of all the chromosomes of an organism's cell, shown in decreasing size of the homologous pairs.

Karyograms can help determine the sex of the organism as well as the possible chromosomal irregularities that might be disease-causing.



One such irregularity in humans is trisomy 21 (an extra chromosome in the 21st pair) that is a cause of the Down syndrome.

23rd pair of human chromosomes determine the sex of a baby.

These chromosomes are called sex chromosomes (as opposed to all other ones that are called autologous).

These chromosomes do not have to be identical; a male individual will have one X and one Y chromosome which are of different sizes, while a female individual will have two X chromosomes of the same size (and banding pattern).

Figure 3.1: Karyogram.



Karyotype is the characteristic pattern of chromosomes of an organism, referring to their size, shape and the banding pattern.

Karyogram is obtained in two possible ways: amniotic fluid sampling and chorionic villus sampling.

Amniotic fluid sampling

A hypodermic needle is inserted through the abdomen of the mother into the amniotic sack.

The embryo swims in the amniotic fluid which contains the cells that the embryo has shed off.

The karyogram is obtained by collecting the DNA from these cells.

Chorionic villus sampling

Similarly, the chorionic villi make up the embryonic side of the placenta.

These villi are of the embryonic tissue origin which means that they contain the same cells as the child.

By sampling the chorionic villi (again using a needle), the child's cells can be obtained and karyogram constructed.

3.2 Meiosis



Meiosis is a type of cell division in which one cell with a diploid nucleus divides into 4 genetically distinct cells with haploid nuclei. This is the process whereby gametes (sex cells) are made.

Diploid cells have two variations of each chromosome (one maternal- and one paternally derived)

Haploid cells have one variation of each chromosome (either a maternal- or paternally derived one)

The AIM of meiosis is to create cells with half the number of chromosomes, so that during fertilization, each parent could contribute their own set of genes to the offspring and thereby conserve the number of chromosomes of a species and promote variation.

Meiosis consists of two divisions: meiosis I (or reduction division); and meiosis II, which can be referred to as mitotic division.

In meiosis I, the number of chromosomes is halved between the two newly formed cells (therefore the name “reduction”) while in meiosis II, the number of chromosome between the parent and daughter cells stays the same, but the chromatids separate (therefore the name mitotic).

The stages of each meiotic division have the same name as in the mitotic division, but the events of each stage differ slightly.

Meiosis also requires duplication of DNA, which occurs prior to first meiotic division.



Homologous pairs are a set of one maternal and one parental chromosome that have the same genes in the same loci along the chromosome. Remember that diploid organisms have two variations of each chromosome number (for example, you have two chromosome 21s)

Sister chromatids are identical copies formed by the replication of a chromosome. These are joined at the centromere in a replicated chromosome. Once sister chromatids separate, they are referred to as **individual chromosomes**

3.2.1 Meiosis I

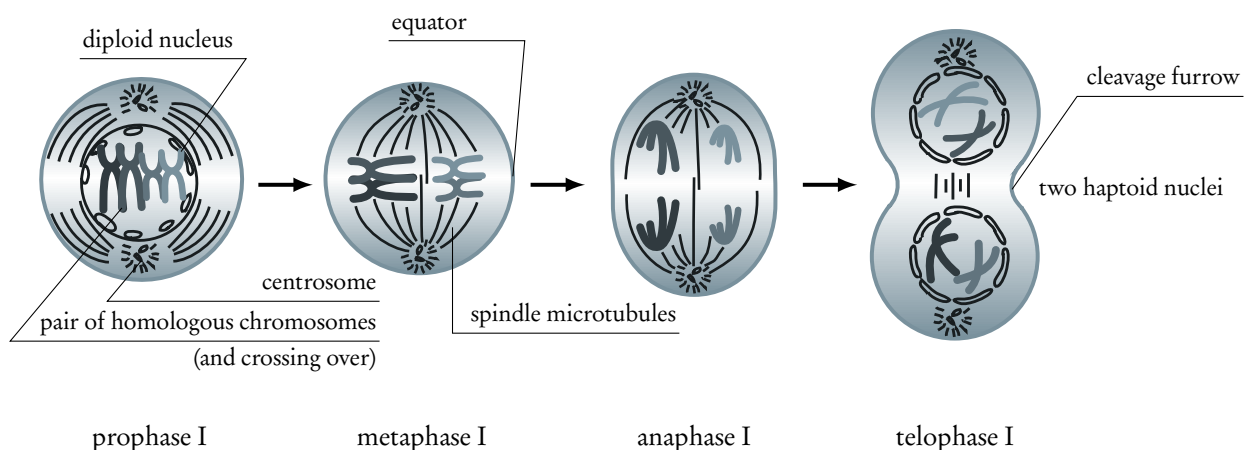


Figure 3.2: Meiosis I.

Prophase I



- Prior to this phase, the DNA has already duplicated (S phase) and the cell contains double the number of chromosomes (depicted as two chromatids of each chromosome).
- DNA supercoils and chromosomes shorten.
- Nuclear envelope breaks down
- Centrioles move to the poles.

Metaphase I



- **Homologous chromosomes pair up** at the cell equator (this means that the pairs of each chromosomes line up on top of each other).
- **Each pole's spindle microtubule attaches to one chromosome from each homologous pair** (recall that in mitosis, one chromosome would have one of each pole's spindle microtubules)

Anaphase I



- **Each spindle microtubule pulls one whole chromosome of the homologous pair towards its pole, causing a division of the chromosome pairs across the cell** (recall that in mitosis, at this step, only the chromatids would separate so that a full set of chromosomes would be present at each pole).
- The movement of the **chromosomes** is achieved through shortening of the spindle microtubules.

Telophase I



- Nuclear envelope forms around each set of chromosomes.
- The cell divides into two cells with **haploid number of nuclei** (only one chromosome from each homologous pair).
- Chromosomes partly uncoil.
- **The cell will proceed with meiosis II**

3.2.2 Meiosis II

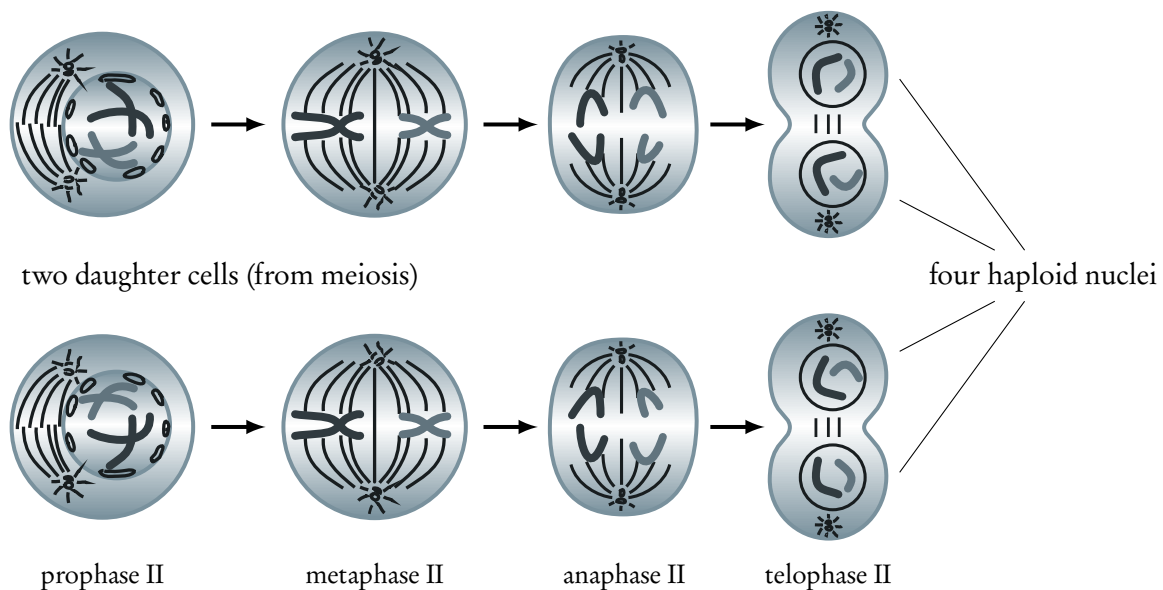


Figure 3.3: Meiosis II.

Before meiosis II, there is no duplication of DNA, so the cell proceeds straight from the telophase into the new division.

Recall that each cell now possesses one chromosome of each homologous pair, but with two chromatids (meaning that each chromosome is made up of two sister chromatids with identical DNA!)

Note that the events of this phase are identical to mitosis, but the starting number of chromosomes is halved!

Prophase II



- Chromosomes supercoil again and become shorter.
- Centrioles again move to the poles of the cell.
- Nuclear envelopes break down.

Metaphase II



- Chromosomes line up at the metaphase plate, one next to each other across the equator.
- Spindle microtubules (one from each pole) attach to the centromeres of the chromosomes (centromeres).

Anaphase II



- Spindle microtubules pull the sister chromatids apart, so that one chromatid of each chromosome travels to the opposite pole.
- Therefore, each pole of the cell will receive one DNA copy of each chromosome.

Telophase II



- At this last stage, each pole of the cell contains half the number of chromosomes compared to beginning of meiosis I, but the same number of chromosomes (just half the chromatids) compared to meiosis II.
- The nuclear envelope forms and the cell divides into two cells.
- Recall that in the first meiotic division, two cells were formed, meaning that now, each of those two cells divided into two, yielding a total of 4 cells, each with half the number of chromosomes

3.2.3 Meiosis and variation

When we talk about variation and meiosis, it is important to understand the following terms:



Crossing over process through which the non-sister chromatids within a homologous pair, exchange genetic material during prophase I.

Through exchange of genetic material between non-sister chromosomes of a homologous pair, gametes end up with chromosomes with new gene combinations that were not present before. These are called recombinants.

During alignment of homologous pairs in prophase one, non-sister chromatids cross over, forming an x-shaped form called chiasmata.

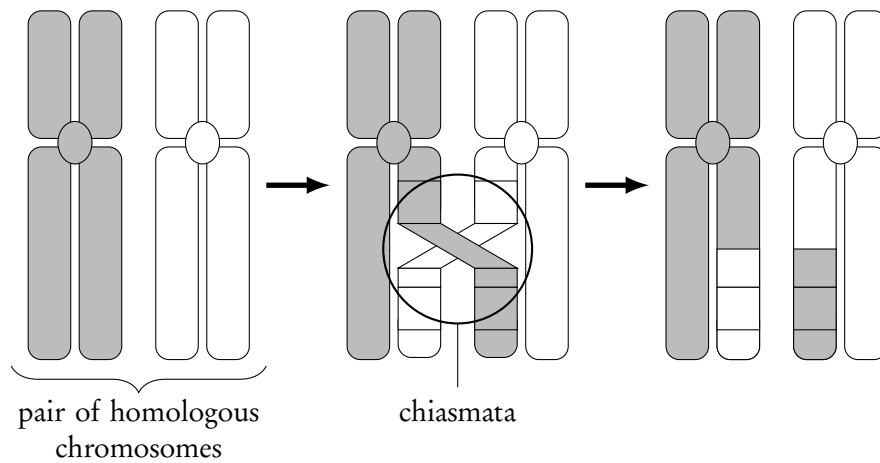


Figure 3.4



Random orientation refers to the fact that the positioning of each homologous chromosome at the metaphase plate during metaphase I is random (not pre-determined).

Recall that, after prophase I, each chromosome in a homologous pair possesses different allele combinations.

Depending on which chromosome orients towards which pole in metaphase I, each cell will end up with a different set of chromosomal alleles.

The orientation of one pair of homologous chromosomes in meiosis I is independent of the orientation of any other pair, so the process is called independent assortment.

If these definitions are clear, then it is obvious how within one individual, there is an infinite variation between the gametes. Now, adding the process of fertilization, where two random gametes from two different individuals of a species fuse, there is truly an infinite number of possible combinations and therefore infinite genetic variation.

3.2.4 Failures of meiosis

Recall that failures of checkpoint mechanisms during mitosis often result in tumour formation. The failures of meiosis, on the other hand, are usually lethal for the embryo, or result in genetic disorders.

Example.

Down Syndrome

Down syndrome is a chromosomal abnormality in which the individual possesses 1 extra chromosome in their 21st chromosomal pair.

This is a result of improper separation of homologous chromosomes during anaphase I, or improper disjunction of sister chromatids during anaphase II.

The cell that ends up with an extra chromosome in the 21st pair will be able to continue its life cycle through fertilization.

The cell that ends up with one less chromosome will not be able to continue its life cycle and will not survive past fertilization.

Disjunctions are believed to be common, but most of them are lethal to the embryo and present themselves as spontaneous abortions.

3.3 Inheritance



Allele a variation of a gene that differs from another allele by a few bases only. This results in variations of a characteristic in an organism. A gene can have none as well as several alleles.

Genotype combination of alleles of one or more genes (literally capital and lower case letters in Punnett grids). Remember that diploid organisms have two copies of each chromosome, which allows two to have two alleles per gene.

Phenotype the physical trait that is expressed by a certain genotype (what you can see with your eyes, like eye colour etc.)

Homozygous two of the same alleles.

Heterozygous two different copies (of an allele).

Dominant allele allele that is expressed both in homozygous and heterozygous combinations.

Recessive allele allele that is only expressed in homozygous combinations.

Co-dominant dominant alleles that are both expressed when present, since neither of them overpowers the other ones.

Locus is a fixed, physical location on a chromosome where genes can be found.

3.3.1 Mendel's law of inheritance

Mendel discovered some basic laws of inheritance by cross-fertilizing pea plants with different traits (flower colour, pea shape). He observed “hidden” traits that tend to surface after several generations. These were in fact what we now call recessive alleles.

Since gametes possess only one set of chromosomes (haploid) compared to somatic cells, they also possess only one allele that they can pass onto the offspring. The other parent will give the other allele of the gene.

Since alleles are variations of traits (e.g., a gene codes for hair colour in general, but the alleles code for the specific hair colours), the combination of alleles will determine the final trait of the individual.

A Punnett grid is a useful tool to predict all the possible offspring combinations of a particular trait.

3.3.2 Punnett grids

Represent the maternal alleles on one side, and paternal alleles on the other side of the grid.

In the first step, based on the parent's full set of chromosomes, one can determine what possible alleles the parents' gametes can have.

In the second step, by combining all the possible parents' alleles in the grid, one can determine the possible offspring combinations.

		maternal alleles	
		A	B
paternal alleles	C	AC	BC
	D	AD	BD

Example.

Rules of Punnett grids

Parental generation is called P1, first offspring generation F0 and all other offspring generations are numbered F1, F2 etc.

Dominant alleles determine the letter used to describe the trait (if brown eyes are dominant to green eyes, then the trait will be defined by letter B and not G).

Dominant alleles are written in the capital letter, and recessive alleles in the lower case letter (therefore, brown eyes will be B, and green eyes will be b, not g!).

Example.

Pea flower colour (purple allele dominates the white allele)

	Parent 1	Parent 2
Genotype	PP (homozygous dominant)	pp (homozygous recessive)
Possible gametes	P and P	p and p
Phenotype	Purple flowers	White flowers

The Punnett grid is constructed by placing the alleles of one parent on the vertical side of a 2×2 square, and the other parent on the horizontal side.

		Parent 1		P1:	PP	pp	
		P	P				
Parent 2	p	Pp	Pp	F0 Genotype	Heterozygous (dominant) Pp		
	p	Pp	Pp	F0 Phenotype	Purple (since one dominant allele is always present)		

Let's see now what happens if you cross-pollinate the offspring.

		Parent 1		P1:	Pp	Pp	
		P	p				
Parent 2	P	PP	Pp	F1 Genotype	25% PP, 25% pp, 50% Pp (25% HomD, 25% HomR, 50% HetD)		
	p	Pp	pp	F1 Phenotype	75% purple, 25% white		

3.3.3 Co-dominant alleles

Recall that co-dominant alleles are those where neither of the alleles over-rules the others. All the present alleles are expressed.

For example: Each individual has one of the four possible blood groups, namely A, B, AB or O. These are in fact the surface molecules carried on the red blood cells (RBCs) that help the body distinguish between self and non-self. An individual may have “A molecules”, “B molecules”, “A and B molecules”, or no molecules (O) on the surface.

Depending on what molecules are expressed in the RBCs of an individual, he/she will have antibodies or an immune reaction to the molecules he/she does not have.

Therefore, for example, a person with A expressed on their RBCs cannot receive blood from someone with B expressed on their RBCs. Additionally everyone can receive from O-type donors as their RBCs express no molecules on the surface. Meanwhile they cannot receive from any donors that are not O, as they recognize anything on the surface as foreign. Overall, it is very dangerous to receive blood from a mismatched donor, as it can lead to one's immune system attacking their own blood.

Alleles that determine the blood groups are either allele for A, for B, or for O. Alleles A and B are co-dominant, and allele for O is recessive to both A and B. Therefore, if an individual has both the allele for A and B, her blood group will be AB, but if she has allele A and allele O, her blood group will be A.

Rules of Punnett grids

- For co-dominant alleles, the letter used to represent all alleles is capital I.
- For the specific alleles, for example in the ABO blood group example, blood group A is labelled as capital I with a superscript A, so I^A and blood group B as I^B .
- If there is another allele that is recessive to both the co-dominant alleles, that one is labelled as a lower case letter i with no superscript.

Example.

ABO blood groups

	Parent 1	Parent 2
Phenotype	Blood group A	Blood group B
Genotype	$I^A i$ (heterozygous A)	$I^B i$ (heterozygous B)
Possible gametes	I^A and i	I^B and i

		Parent 1		P1:	$I^A i$		$I^B i$			
		I^A	i							
Parent 2	I^B	$I^A I^B$	$i I^B$	F1 Genotype	25% $I^A i$, 25% $I^B i$, 25% $I^A I^B$, 25% ii					
	i	$I^A i$	ii	F1 Phenotype	25% A, 25% B, 25% AB and 25% O					

Example.

Another example

	Parent 1	Parent 2
Phenotype	Blood group AB	Blood group O
Genotype	$I^A I^B$ (co-dominant)	ii (homozygous recessive)
Possible gametes	I^A and I^B	i and i

		Parent 1		P1:	$I^A i$		$I^B i$	
		I^A	I^B					
Parent 2	i	$I^A i$	$I^B i$	F1 Genotype	50% $I^A i$, 50% $I^B i$			
	i	$I^A i$	$I^B i$	F1 Phenotype	50% A and 50% B			

3.3.4 Sex linkage

Recall that the chromosomes can be divided into somatic and sex chromosomes. The sex chromosomes are the ones that determine the sex characteristics of an organism, amongst others.

Sex chromosomes are labelled as X and Y. Y chromosome is much smaller: for that reason, X chromosome carries more genes than the Y chromosome does.

The presence of Y chromosome determines that gender of the child will be male: females normally carry two X chromosomes, while males carry one X and one Y chromosome.

This means that the female will pass on only X chromosomes to her offspring, while the male can pass on either an X or a Y chromosome.

In order for a female offspring to be born, gametes carrying an X in both male and female cell must meet, and for a male to be born, a female gamete with X and a male gamete with Y have to fertilize.



Sex linkage refers to the inheritance of genes that are located on the sex chromosomes.

Since the X chromosome carries more genes than the Y chromosomes, males will often lack one copy (and therefore one allele) of the sex linked genes. For this reason, many of the sex linked diseases affect males to a higher degree than females.

Rules of Punnett grids

- For sex linked genes, the letters assigned to the traits are either X or a Y, depending on the gender of the individual.
- The trait is labelled as a superscript on the X or Y.
- Dominant allele is written in the capital letter, while the recessive is written in the lower case letter.

Red-green colour-blindness

Red-green colour-blindness is a sex linked disorder carried on the X chromosome where the affected individuals cannot distinguish between red and green colours. It is a recessive disorder, meaning that only individuals with no dominant (healthy) alleles are affected by it. However, since it is a sex linked trait, carried on the X chromosome, males already have a disadvantage since they have only one chromosome that can either carry the healthy or the affected gene.

Females carry two X chromosomes, and therefore two alleles for colour vision. Males carry one X chromosome with the gene for colour vision, and a Y that doesn't contain the gene at all. If a female has one healthy and one affected gene, she will be called a carrier but she will be healthy. She might pass on her affected X chromosome to her son, who will not inherit another X chromosome, but a Y chromosome from his father and therefore be affected by the disease. Note that females can be affected as well, but in that case they would have to have a carrier mother and an affected father (and still there is only 50% chance that they will have the disease).

Let's look at what happens in the Punnett grid.

Parent	Mother	Father
Phenotype	Healthy (carrier)	Healthy
Genotype	$X^H X^h$	$X^H Y$
Possible gametes	X^H and X^h	X^H and Y

		Parent 1		P1:	$I^A i$	$I^B i$
Parent 2		X^H	X^h	F1 Genotype	25% $X^H X^H$, 25% $X^H X^h$, 25% $X^H Y$, 25% $X^h Y$	
	X^H	$X^H X^H$	$X^h X^H$	F1 Phenotype	75% healthy child, 25% affected child (but 50% that the son will be affected, and 0% chance that the daughter will be affected)	
	Y	$X^H Y$	$X^h Y$			

3.3.5 Dominant and recessive genetic disorders

So far, the traits we have shown have all been recessive. This means that only the individuals with two affected alleles get the disorder. If the disease is of dominant inheritance, one affected allele is enough for the individual to carry the disease. This will result in a much higher percentage of affected individuals in a family tree (compared to the recessive disorders).

Example.

Huntington's disease

	Parent 1	Parent 2
Phenotype	Healthy	Huntington's
Genotype	Hh (homozygous recessive)	Hh (heterozygous dominant)
Possible gametes	h and h	H and h

		Parent 1		P1:	I ^A i	I ^B i
		h	h			
Parent 2	H	hH	hH	F1 Genotype	50% Hh and 50% hh	
	h	hh	hh	F1 Phenotype	50% affected offspring, 50% healthy offspring	

3.3.6 Pedigree charts

Pedigree charts are a way to represent the inheritance of certain traits in a form of a family tree, where the oldest individuals are set at the top, and their offspring follow downwards.

There are some rules you should keep in mind:

- The female is always labelled as a circle ○ and the male as a square □.
- Most pedigree charts show affected individuals as coloured figures ●■ and healthy and carrier individuals as transparent figures ○□.
- Such a chart helps determine the possible genotypes of individuals and chances for affected offspring in the future.

Note that in the exam, you could be asked to also determine whether the disease is sex linked, recessive or dominant.

Here are again some tips that might help you out with that

- Charts where mostly males are affected usually represent a sex-linked trait (this means that more than 90% of the affected individuals are males).

- Two healthy individuals cannot have a child with a dominant disorder.
- In recessive disorders, two healthy parents can have an affected child, but two affected parents cannot have a healthy offspring.
- Try to annotate the diagrams as much as possible to help you keep track of what you figure out.

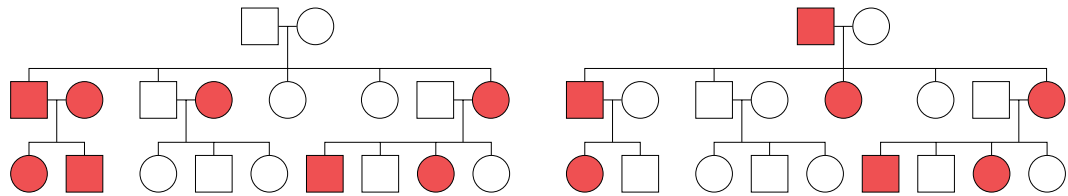


Figure 3.5: Recessive vs. dominant chart.

3.3.7 Dihybrid crosses

In a monohybrid cross, we only looked at the inheritance of one trait on one chromosome. In a dihybrid cross, two traits can be followed in a Punnett grid to predict the possible combinations of trait.

In a typical dihybrid cross, we assume that each trait is located on its own chromosome, and that they are inherited separately. A typical example involves observing the shape of a pea seed and the colour of its flowers.

Example.

Pea seed shape and colour of flowers. Round seeds are dominant to wrinkled and purple flowers are dominant to white ones.

	Parent 1	Parent 2
Phenotype	Purple flower, round seed	Purple flower, round seed
Genotype	PpRr (heterozygous for both traits)	PpRr (heterozygous for both traits)
Possible gametes	PR, Pr, pR, pr*	PR, Pr, pR, pr*

Note that if we assume that each trait is on a different chromosome, due to independent assortment of chromosomes, each allele of one trait can end up in the combination with any of the alleles of the other trait.

Example.

Punnet grid

		Parent 1			
		PR	Pr	pR	pr
Parent 2	PR	PPRR (R)	PPRr (R)	PpRR (R)	PpRr (R)
	Pr	PPRr (R)	PPrr	PpRr (R)	Pprr
	pR	PpRR (R)	PpRr (R)	ppRR (R)	ppRr (R)
	pr	PpRr (R)	Pprr	PpRr (R)	pprr

P1:	PpRr	PpRr
F1 Genotype	PPRR, PPRr, PPrr, PpRR, ppRR, PpRr, Pprr, pprr	
F1 Phenotype	9:3:3:1 9 Purple round, 3 Purple wrinkled, 3 white wrinkled, 1 white wrinkled	

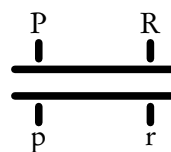
The typical 9:3:3:1 ratio is often observed in a dihybrid cross of non-linked genes (traits in different chromosomes), involving to heterozygous parents for both traits. This ratio is disturbed if the parents are not heterozygous for both traits, if some of the alleles are co-dominant or if the genes are sex-linked.

3.3.8 Linked genes

Recall that sex-linked genes were located on the sex chromosome. The term “linked-genes” refers to genes that are located on the same autosomal (non-sex) chromosome. In the previous example, the genes were not linked and the allele combinations were inherited separately. This is why all combinations of alleles were possible.

In linked genes, two genes are on the same chromosome, which means that the alleles of those two genes are inherited together.

When genes are linked, they are graphically represented as follows:



This suggests that alleles PR and alleles pr will always be transmitted together, and even though this parent is heterozygous for both genes, it will only give gametes with two possible combinations.

Since each chromosome contains many genes, the combinations of alleles of these genes would always be same, which would lead to very little variation between individuals. However, recall that in the process of meiosis, we mentioned a term called “crossing over” that introduced extra variation between individuals.

3.3.9 Crossing over and linked genes

If there were no crossing over, a cross between two linked genes, of two heterozygous parents could either donate PR or pr chromosomes.

However, during crossing over, there is a sharing of genetic material, depicted in Figure 3.6.

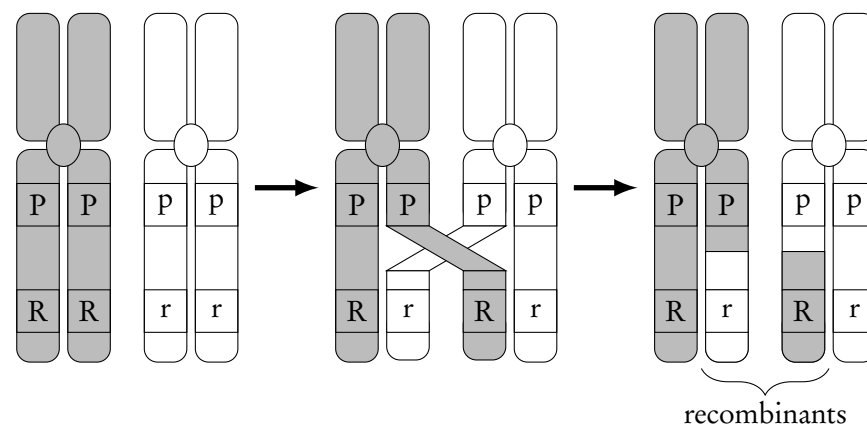


Figure 3.6

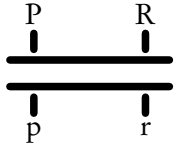
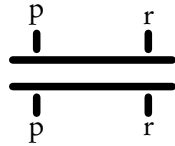
Therefore, the new parental gamete combinations are PR and pr (the combinations from the original chromosomes), and Pr and pR (the recombinants, or new combinations that did not exist in the parent chromosomes). However, as you can imagine, crossing over does not happen in the same locus in every gamete, so, in a cross of a thousand gametes, the two original combinations, PR and pr will be by far more common than the recombinants. For this reason, the majority of children will have the original parental combinations (PR and pr), while a minority will have the recombinant genes (pR and Pr).

Possible exam questions

Keep in mind that the further the two genes are in the chromosome, the more chances there are for crossing over to separate the alleles and create recombinants.

Example.

Identify the possible F2 genotypes in a cross of a heterozygous purple, round pea and a homozygous white wrinkled pea.

F1:	PpRr	prrr
		
Gametes:	PR & pr + Pr & pR (after crossing over)	pr
F2:	PpRr and prrr (majority) + prrr and pPRr (minority)	

3.3.10 Polygenic inheritance

Polygenic inheritance refers to the phenomenon where one trait is controlled by multiple genes. When a trait is controlled by more than one gene, the result is often a wide range of variations between the individuals.

Discrete variation occurs when a trait is controlled by one or only a few genes. In traits that show discrete variations, individuals either have the trait or don't. An example of this could be Huntington's disease, where you either carry the dominant allele for the disease, and you have it, or you don't, and you're healthy.

In continuous variation, the trait is controlled by many genes, and they are often co-dominant. In such a case, the genes have a cumulative effect. An example of such variation is skin colour, or height.

Example.

Polygenic inheritance of skin colour

Skin colour is controlled by multiple genes, each determining the presence of the melanin pigment (the pigment that gives skin a dark colour) in the skin cell. Each gene has two possible alleles, expression of melanin, or no melanin expression. If an individual has a set of genes where the majority have the allele for melanin expression, the colour of that person's skin will be dark.

The more the alleles for melanin production, the more the pigment this person will have in their cells. If many genes control a certain characteristic, the distribution of such characteristic will be close to the normal distribution.

In the example of human height, many genes control height, so there is a continuous variation of human heights. Still, most people will be of average height, and some people will be very tall or very short.

Example

Genes A, B, C and D code for melanin production. They come with two alleles, D (dark) and L (light), which are co-dominant.

A person with $A^D A^D B^D B^L C^D C^D D^D D^L$ will have a darker skin colour compared to a person with $A^L A^L B^L B^L C^L C^L D^D D^D$.

3.4 Genetic modifications and biotechnology

3.4.1 PCR and gel electrophoresis

A single cell contains a miniature amount of DNA. A DNA sample, obtained from a piece of tissue or bodily fluid is often not enough on its own to give conclusive results. For that reason, the DNA needs to be multiplied to make sure there is enough for proper analysis:

Polymerase chain reaction (PCR)

PCR is a method through which a molecule of DNA is copied many times, yielding a high number of identical DNA molecules. The system is based on the cellular replication process but employs the use of the polymerase enzyme from *Thermus aquaticus*, which works best at high temperatures. At high temperatures, the two strands of DNA molecules naturally separate, and the polymerase enzyme is able to replicate both chains. When this process finishes, the temperature is lowered, so that the chains join again. This is repeated through several cycles, until the DNA is sufficiently quantified. You can imagine this goes quite fast, since from one DNA molecule, you get two, which can then be amplified to another four, then 16 and so on.

In synthesis, the steps of PCR are:

- Denaturation: ($\sim 95^\circ\text{C}$) the DNA sample is heated to separate the double stranded molecule into two single strands.
- Annealing: ($\sim 55^\circ\text{C}$) the DNA primers attach to the 3' ends of the target sequence.
- Elongation: ($\sim 72^\circ\text{C}$) the heat-tolerant DNA polymerase (Taq) binds to the primers and copies the DNA strands (e.g., from 2 to 4 strands).

These steps are repeated over and over until a large number of copies from the original molecule is obtained.

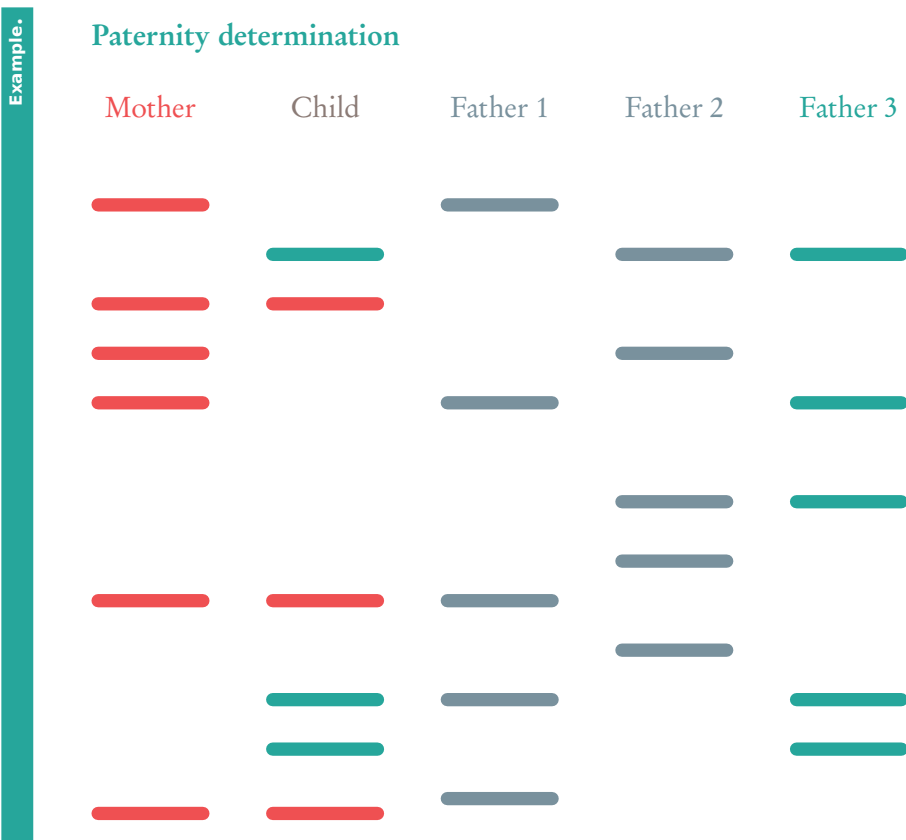
Gel electrophoresis

This method was developed to visualize the DNA fragments, or proteins isolated from cells. The principle of this method is to separate DNA fragments or proteins from a mixture, based on their size and charge. The electrophoresis chamber consists of an agarose gel submerged into a liquid, and two electrodes (a positive and negative) at each side. The voltage across the chamber makes the molecules of different sizes and charges move across the gel at different speed. Large and less charged molecules stick behind, while the small and highly charged molecules travel faster.

The result is a pattern of bands visualized under UV light which can be compared between cells, or individuals (in case of paternity tests).

Gel electrophoresis can be used to determine the paternity of a child by comparing DNA fragments of the mother, the child and the suspected fathers.

Since the child's DNA is a combination of his father's and mother's DNA, the child's pattern will contain some of his mother's and some of his father's bands



3.4.2 Genetic modifications

Recall that the genetic code is universal, meaning that all the organisms translate the same triplets of bases into the same amino acids. With that in mind, you can see why a gene in one organism's coding for a specific protein, could be transferred to another organism's where it would again code for the same protein.

Example.

E. coli and the production of insulin

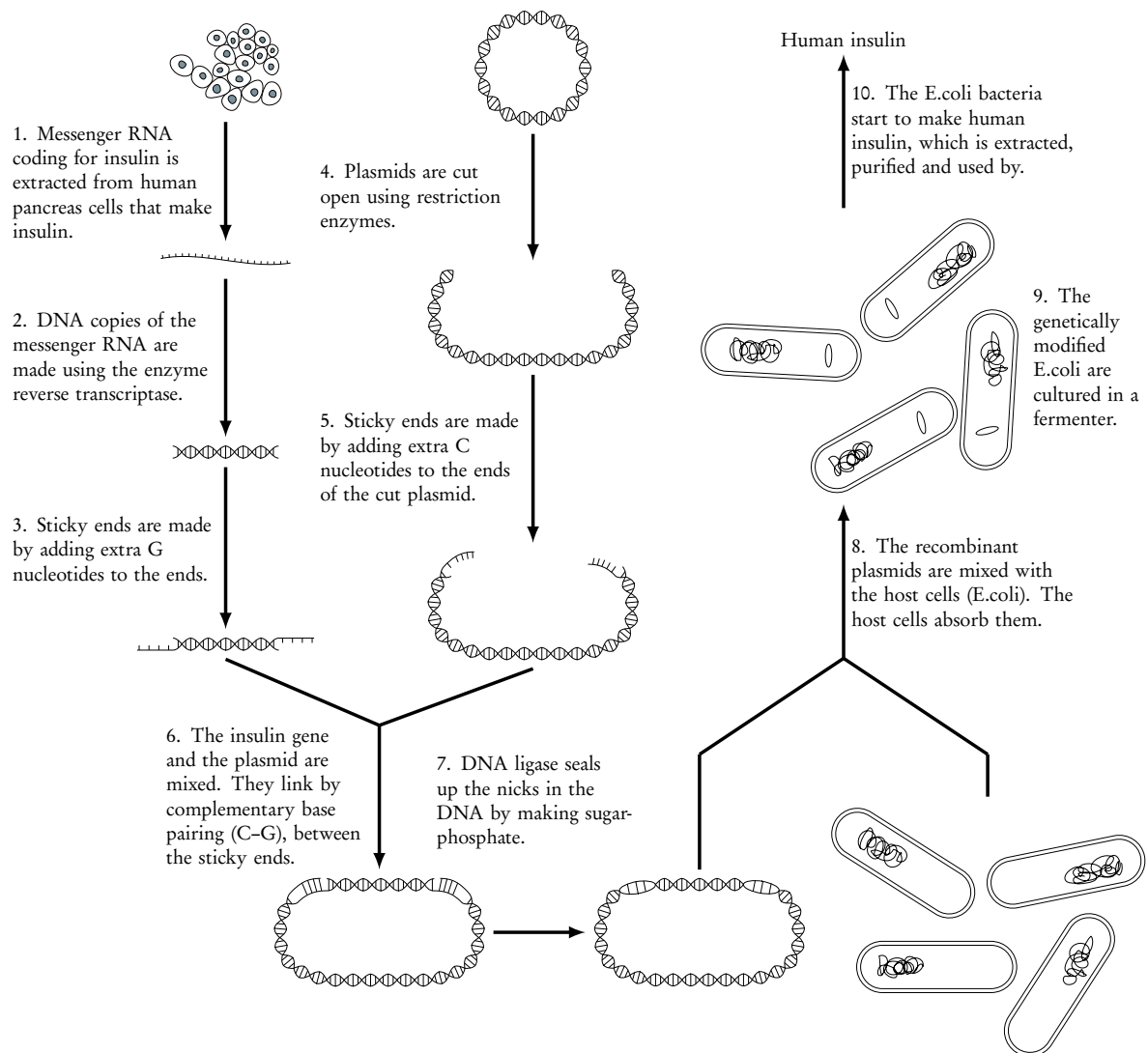
Bacteria can be genetically modified to produce a human protein of interest. An example of this is the production of insulin through E. coli. This is achieved by extracting the messenger RNA from human cells producing insulin (pancreatic cells) and converting that mRNA back into DNA using reverse transcriptase enzyme.

This piece of DNA could now be ligated into a plasmid using restriction enzymes, enzymes that are able to cut DNA at a specific point with a specific pattern, leaving sticky ends through which the gene of interest (from previous step) could be ligated into the plasmid (which has also been cut with the same restriction enzymes).

Once the plasmid with the gene of interest is made, it can be incubated with the bacteria which will then take it up and start transcribing and translating the newly acquired gene. These bacteria are now called recombinant bacteria.

The result will be the production of the protein, which can then be collected as a part of the bacterial secretions.

Example.



3.4.3 Benefits and risks of genetic modifications

Benefits

- With less pest damage, there are higher crop yields, so less food shortages.
- With better yield, less land is needed for the same amount of crop production, and the unused land can be conserved for wildlife.
- There is no need to use pesticides that damage other organisms living in the vicinity.

Risks

- Long term effects of genetically modified foods have not yet been determined.
- The pollen of the modified crops might be blown away to the wildlife where it might kill the organisms that do not normally infect the plant.
- The genetically modified plants have an evolutionary advantage to the non-modified plants, so the random cross-pollination might create an imbalance in the ecosystem.

Example.

Bt maize

Bt maize can be genetically modified to express a bacterial gene for Bt toxin that is toxic to the pests that usually attack it. In that way, the crops are “resistant” to the pest infestations.

3.4.4 Clones and cloning



Clone an organism that is genetically identical to its parent organism.

Cloning is a technique of producing genetically identical cells, tissue or organisms. This is usually done to obtain a higher number of cells or individuals with a desirable set of characteristics. Cloning of plants can be relatively easy by taking a piece of root or stem that contains plant stem cells that will grow into a new, genetically identical plant. Animal cloning is more difficult since animals cannot develop from a group of stem cells found in the body.

Dolly the sheep

Dolly the sheep was the first cloned animal. Three sheep were used to give rise to Dolly, namely one that donated the egg cell (without the nucleus), one sheep that donated its genetic material (nucleus of a somatic cell) and one sheep into which the embryo was implanted. Therefore, Dolly was genetically identical only to the sheep donating the nucleus.

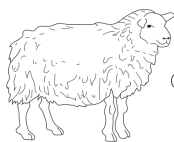
Process

An egg cell was obtained from a sheep, and the nucleus of that cell was removed. The egg cell contains only half the genetic information, so its nucleus cannot be used in cloning. However, the egg cell contains the vital enzymes and organelles for development.

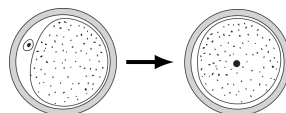
Udder cells (somatic cells) from another sheep were taken and grown in deprived environment which cause them to switch off all their genes. A nucleus of such a cell was isolated (it contained all the chromosomes), and then fused with the egg cell without the nucleus.

The fused cells were inserted into the third sheep (the surrogate) and they developed as a newly formed embryo. The sheep that was born was named Dolly and was genetically identical to the mother providing the nucleus of its cell.

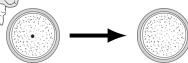
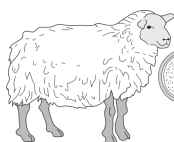
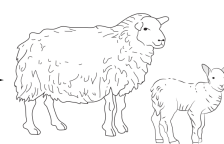
Cell taken from udder of donor adult and cultured in laboratory for six days



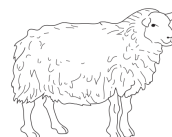
Egg without a nucleus fused with donor cell using a pulse of electricity



Surrogate mother gives birth to lamb. Dolly is genetically identical with the sheep that donated the udder cell (the donor)



Embryo resulting from fusion of udder cell and egg transferred to the uterus of a third sheep which acts as surrogate mother



Unfertilized egg taken from another sheep. Nucleus removed from the cell

3.5 Gene pools and speciation



Species is defined as a group of organisms that can interbreed and give fertile offspring.

Population is a group of individuals of the same species living at the same area at the same time.

Gene pool refers to all the possible genes and their alleles within a species.

Evolution is defined as gradual change in the heritable characteristics of an individual.

This change can be caused by genetic mutations which can be either beneficial or detrimental for the species.

Evolution can also be a result of environmental pressure that favour one characteristic within the species over others.

Speciation refers to the process where a new species is formed from a pre-existing species.

Keeping these definitions in mind, let us understand how evolution is determined by natural selection and therefore the shifts in allele frequencies.

3.5.1 Directional, stabilizing and disruptive selection

Directional

Selection that favours individuals on one end of the phenotypic distribution. Individuals on the other end are not favoured by the environment so are slowly dying out.

An example of this could be the increase in dark moths during industrial revolution, when the pollution caused darkening of the trees (from the smog), so the white moths were more visible to the predators.

Stabilizing

Selection that favours individuals in the middle of the phenotypic spectrum. Both extremes in the spectrum of a characteristic are not favoured by the environment.

Recall the example of sickle cell anaemia. Individuals that are homozygous either for healthy cells, or sickle cells will be exterminated in the areas exposed to malaria. The heterozygous individuals will have both resistance to malaria and a decent oxygen carrying capacity to be favoured by the environment

Disruptive

Selection that eliminates individuals with the intermediate characteristics. In this selection, both ends of the phenotypic variation will be favoured by the environment, but not the intermediate individuals.

Squirrels with long tails are good at balancing themselves on the trees, but the squirrels with short tails are faster at running away from predators on the ground. The squirrels that have intermediate tails are easily caught by the predator on the ground, but also but also on the trees, as their balance is not as good as the really long tailed squirrels.

Speciation can occur thorough reproductive isolation of populations. Three types of reproductive isolation contribute to this:

Temporal: Temporal isolation occurs when two populations stop interbreeding because of differences in their mating times. An example of this could be the maggot fly, which in North America used to lay its eggs on the hawthorn fruits, which were the food for its larvae. With the introduction of non-native apple trees, it started laying eggs on those as well. However, since the two fruits ripen at different times, depending on which fruit the fly lays its eggs, it will mate at different times. Therefore within the population, there is a temporal mating barrier.

Behavioural: Behavioural isolation can occur if the mating behaviour of individuals of the same population starts differing. An example of this was the speciation of fireflies which mate by sending specific light signals. With time, some fireflies started signalling through a different pattern, and only certain individuals would respond to that pattern. Now, there are many different firefly species which do not interbreed because their courtship mechanisms are different.

Geographical: In this type of isolation, individuals of the same population become separated geographically and can therefore not mate anymore. An example of this are the lava lizards from Galapagos. A group of lizards managed to get from one island to another, but as they couldn't return anymore, they could only interbreed with the population that ended up at the island. Over time, the interbreeding of the lizards on one, and on the other island, created two different species of lizards.

Gradual and abrupt speciation

There are full fossil records showing the gradual change of species over time. Environmental pressures and species adaptation would eventually lead to accumulation of new characteristics that differ one older species from a newer one.

However, there are incomplete fossil records where the lack of intermediate fossils could be explained by a more abrupt evolution. This could happen through genetic mutation, like doubling of the chromosome numbers that results in organisms that are incompatible with their parental species.

Polyploidy is an example of such an abrupt change

Polyploidy

Individuals of the same species have the same number of chromosomes. Non-disjunction of chromosomes during meiosis may result in gametes with double the number of chromosomes. If such a gamete is fused with a normal, haploid gamete, the offspring become triploid, and therefore infertile since they cannot divide their chromosomes in half.

This results in a new species, since its individuals cannot mate with other members of the species and produce fertile offspring.

4.1. Species, communities and ecosystems **94**

- Key terms and definitions
- Autotrophs vs. heterotrophs
- Heterotrophs: consumers vs. detritivores vs. saprotrophs
- Nutrient cycling

4.2. Energy flow **97**

- Sunlight: main source of energy in ecosystems
- Food chains and energy pyramids: analyzing energy flow in ecosystems
- Energy loss: limiting food chain length

4.3. Carbon cycling **99**

- Carbon: how does it enter ecosystems?
- The carbon cycle
- Decomposition of organic compounds

4.4. Climate change **101**

- The greenhouse effects and greenhouse gases
- The enhanced greenhouse effect and its effect on global temperatures
- Changes in atmospheric CO₂ concentrations and average global temperature

4.1 Species, communities and ecosystems

4.1.1 Key terms and definitions



Species a group of organisms that can interbreed and produce fertile offspring. Its members can be reproductively isolated (live in a different habitat) but as long as they hold the ability to potentially interbreed with other members from different habitats, they are part of the same species.

Habitat the environment in which a species normally lives, or the location of a living organism.

Population a group of organisms of the same species who live in the same area at the same time.

Community a group of populations living and interacting with each other in the same geographical area.

Ecosystem community and its abiotic environment.

Ecology the study of the relationship between living organisms and between organisms and their environment.

4.1.2 Autotrophs vs. heterotrophs

One way organisms can be classified into different groups is based on their method of nutrition, in other words, by the ways in which they obtain organic compounds necessary to provide essential nutrients to each organism. In this way, organisms can be classified as autotrophs and heterotrophs.



Autotrophs can synthesize their own organic compounds from simple inorganic compounds found in the abiotic environment, to do this, they absorb carbon dioxide, water and other inorganic nutrients from the environment for synthesis (e.g., plants, algae, cyanobacteria (all of these are photosynthetic organisms)). Autotrophs are commonly referred to as producers.

Heterotrophs obtain organic compounds from other living organisms (e.g., humans, insects, fish, etc.). Heterotrophs are commonly referred to as consumers.

Some species can obtain nutrients through both methods (e.g., some bacteria) but this is a rare occurrence.

4.1.3 Heterotrophs: consumers vs. detritivores vs. saprotrophs

Heterotrophs can be divided into three subcategories:

Consumers: organisms that obtain organic matter from other living or recently killed organisms

- Primary consumer: eats autotrophic organisms → herbivore
- Secondary consumer: eats herbivores → carnivore
- Tertiary consumer: eats secondary consumer → top carnivore
(top carnivore can also be a quaternary consumer, depending on the size of the food chain)

Detritivores: decomposers that internally digest dead organic matter like leaves and carcasses (e.g., earthworms and woodlice)

Saprotrophs: decomposers that live in or on dead organic matter and externally digest it, by secreting digestive enzymes and absorbing the products of digestion (e.g., fungus and bacteria)

4.1.4 Nutrient cycling

Since autotrophs sustain themselves by absorbing nutrients and other inorganic compounds from the abiotic environment, there has to be a way in which this inorganic compound supply is maintained. This is achieved through a process known as nutrient cycling. When an organism dies, decomposers are in charge of breaking it down and absorbing as many nutrients and as much energy as possible via digestion. The remains of the organism are then mostly inorganic compounds that return to the abiotic environment after the process of decomposition. Gas balance (oxygen-carbon dioxide) is usually replenished by the processes of cell respiration and photosynthesis in plants and consumers. This returns balance to the ecosystem, once again providing available inorganic matter for autotrophs to make use of. This results in ecosystems having the potential to be sustainable for long periods of time, as long as a source of energy and the necessary fuel sources are available.

Example.

Mesocosms: proving that ecosystems are self-sustainable

The picture below shows one possible design of a mesocosm; an experimental setup that tries to emulate a real-life ecosystem in an isolated space to determine if the ecosystem is indeed self-sustainable:

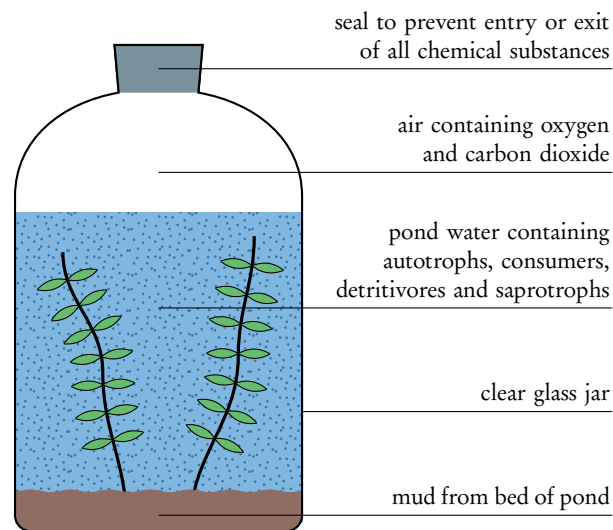


Figure 4.1: Mesocosms proving that ecosystems are self sustainable.

The setup contains the main elements found in ecosystems: a variety of autotrophs and different types of heterotrophs (including decomposers that enable nutrient cycling), an abiotic environment (mud as a source of inorganic compounds and nutrients, water, air containing oxygen and carbon dioxide. With a source of energy (light), this sealed mini-ecosystem should be able to self-maintain for a long time.

4.2 Energy flow

4.2.1 Sunlight: main source of energy in ecosystems

In any given food chain, the first and most abundant level of organisms comprises autotrophs (e.g., plants and algae), also known as producers. Since autotrophs are mainly photosynthetic organisms, they require sunlight as a main source of energy in order to convert it into organic compounds and obtain the necessary nutrients for survival. Without this essential energy source, producers would not survive and primary consumers would not be able to feed, severely disrupting the food chain.

4.2.2 Food chains and energy pyramids: analyzing energy flow in ecosystems

A food chain is a sequence showing the feeding relationships and energy flow between species sharing a habitat. The arrow shows the direction of the energy flow (energy is taken up upon ingestion by the next organism in the food chain).

Example.

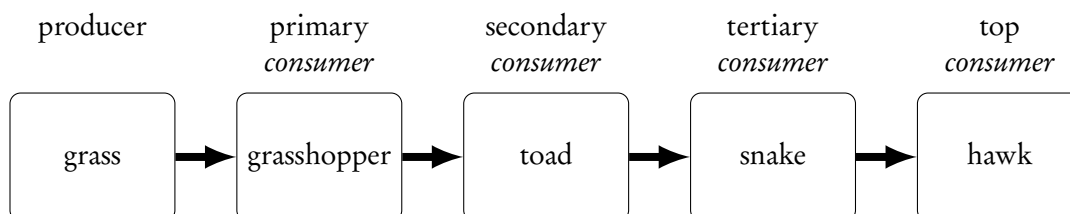


Figure 4.2: Food chains.

A series of interconnected food chains creates a food web, these are much more complicated to draw and interpret (some organisms can be different types of consumers for example). An organism's position in a food chain or web is also known as trophic level, and it is an easy way to classify organisms, the names of each trophic level are shown in grey above the food chain boxes. Energy in the form of carbon compounds flows from one organism to the next when the organism in the higher trophic level feeds on another. However, this energy flow is not 100% effective. We will see why in the next section.

4.2.3 Energy loss: limiting food chain length

Energy enters the ecosystem as sunlight, and is absorbed with about 20% efficacy by autotrophic producers and transformed into chemical energy

Chemical energy obtained from photosynthesis is transferred from one trophic level to the next by means of feeding. The chemical energy stored in bonds is released and made available to an organism when carbohydrates, lipids, proteins and nucleic acids are digested

At each step, only about 10% of the energy is successfully transferred to the feeding organism. This is because energy is lost in several ways:

- Energy released from carbon compounds is used in cell respiration and lost as heat
- When feeding, not the entire organism is eaten (bone, cartilage, etc. is often left to decompose)
- Loss of carbon dioxide, water and other waste products throughout the organism's life
- Some organisms die and decay before being eaten
- Warm-blooded and moving organisms lose more energy as heat

While nutrients can be recycled by decomposers, once energy leaves a food chain it cannot be taken up again

Energy losses of this kind restrict length of food chains and the biomass of the higher trophic levels

A pyramid of energy (example shown in Figure 4.3) shows the flow of energy from one trophic level to another in a community

- The first step is always the largest (producers)
- At each increasing trophic level, the bar becomes 10 times smaller (scaling is important here), as only 10% of energy is transferred
- Energy flow is expressed in units: $\text{kJ/m}^2/\text{year}$ (kilojoules per square meter per year)

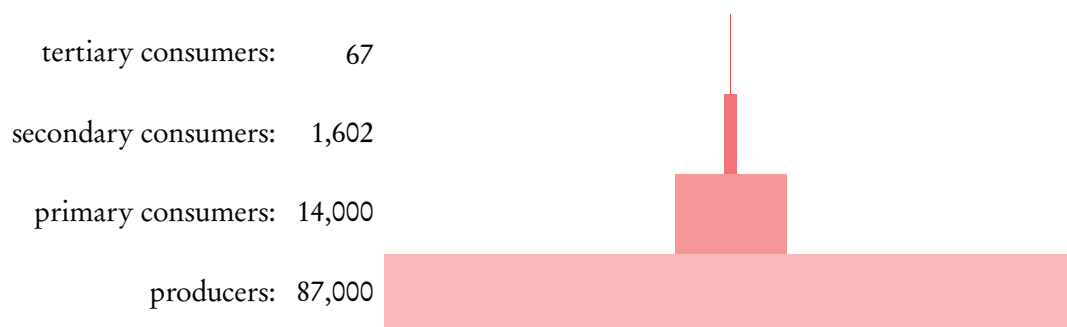


Figure 4.3: Food chains.

4.3 Carbon cycling

4.3.1 Carbon: how does it enter ecosystems?

Carbon often enters an ecosystem in the form of carbon dioxide, an inorganic molecule that diffuses into autotrophic producers.

Autotrophs then have the ability to convert carbon dioxide into complex carbohydrates (organic molecules) and other carbon-containing molecules (e.g., proteins and lipids) through the process of photosynthesis.

Carbon dioxide is present in air; so terrestrial autotrophs can directly absorb it through specialized porous cuticles.

In aquatic ecosystems however, carbon can be found as dissolved CO_2 or in the form of HCO_3^- (hydrogen carbonate ions), which result from the chemical reaction when water and carbon dioxide combine to form carbonic acid (H_2CO_3).

These molecules can then be used in photosynthesis to produce organic compounds. These organic compounds are then used in the process of cell respiration, leading to the regeneration of carbon dioxide, which is released into the water or the atmosphere as a by-product.

4.3.2 The carbon cycle

As stated earlier, carbon dioxide enters ecosystems by diffusing into autotrophs, and is transformed into organic compounds through photosynthesis. Some carbon dioxide returns to the atmosphere as a by-product of cell respiration that takes place in producers. Primary consumers feed from producers, taking up some carbon-containing compounds (and secondary consumers feed on primary, etc.).

Cell respiration by all living organisms leads to further generation of carbon dioxide that is released into the atmosphere.

When consumers die, the leftover carbon molecules in their bodies are acquired by decomposers who eat them.

- As organic matter gets buried into the ground, it can end up in areas with no, or little, oxygen. Here, archaeans called methanogens produce methane from the metabolic by-products of anaerobic respiration.
- The methane will either diffuse into the atmosphere or accumulate under ground, forming natural gas deposits.

- Methane that diffuses into the atmosphere affects carbon dioxide concentrations as it then gets oxidized into water vapour and carbon dioxide,. However, methane persists for about 12 years in the atmosphere before being naturally oxidized.
- The rising number of cattle nowadays has also dramatically increased the methane concentrations released into the atmosphere.

Partial decomposition of dead organisms and fossilization lead to the formation of fossil fuels (more details in the next section)

Combustion of fossil fuels in industrial factories produces carbon dioxide that is released into the atmosphere in great quantities

All of the aforementioned interactions create a cycle of movement of carbon dioxide between the spheres of the earth. These include: the atmosphere (air), lithosphere (ground), hydrosphere (water), and biosphere (living organisms).

Skill: Construct a diagram of the carbon cycle.

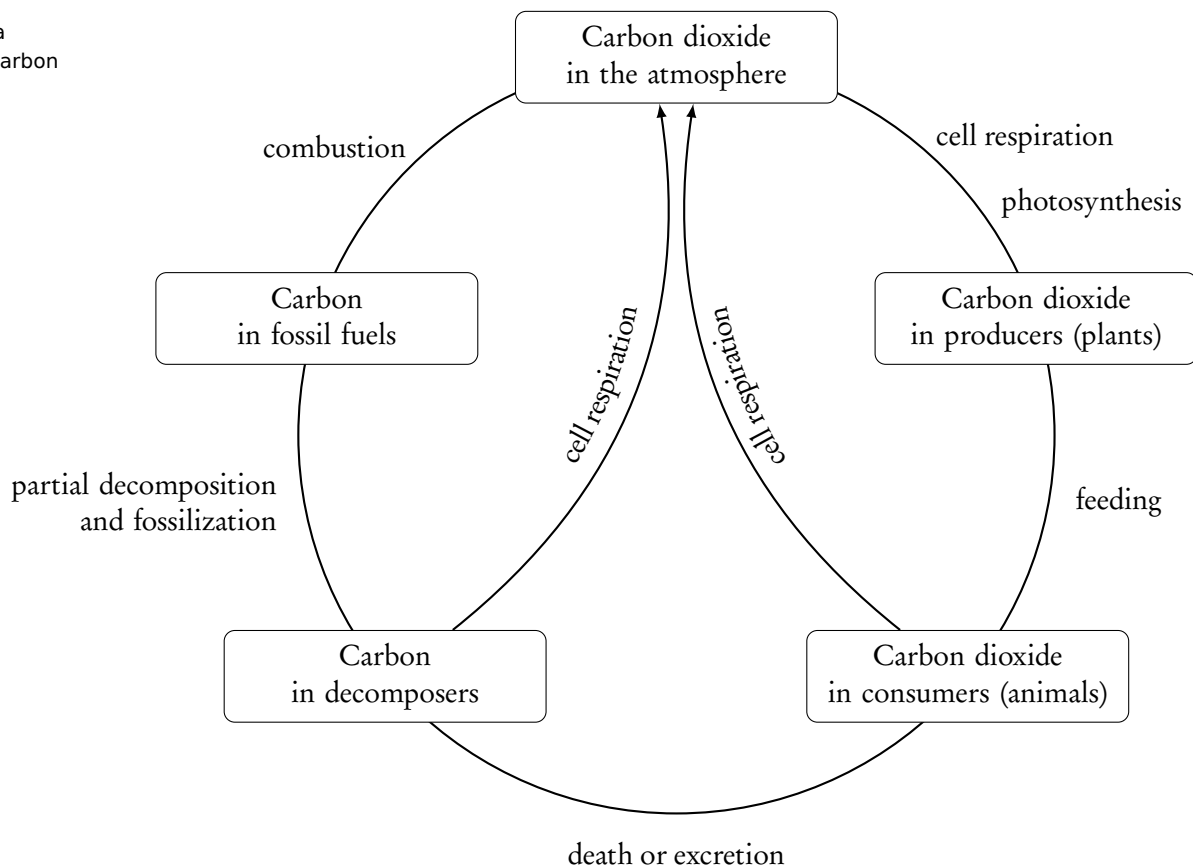


Figure 4.4: Carbon cycle.

4.3.3 Decomposition of organic compounds

Formation of peat and coal

In acidic and anaerobic conditions (e.g., swamps and bogs), saprotrophs cannot fully break down dead organic matter as they require oxygen to function properly.

Partially decomposed organic matter accumulates to form thick deposits called peat.

Peat deposits get compressed under sediments, and with increased heat underground, it forms into a highly concentrated material that undergoes chemical reactions and becomes coal.

Formation of oil and gas

Silt and the remains of dead organic organisms can sometimes accumulate in shallow seas.

Due to the anaerobic conditions, these organisms are only partially decomposed.

Like in peat formation, the silt accumulates and is buried under layers of sediment over time. Through this compaction and heating, hydrocarbons are formed, which make oil and gas, a process that takes millions of years.

The oil and gas formed is forced out of the source and it accumulates in porous rocks. It is then forced out of there when humans drill into such rocks.

4.4 Climate change

4.4.1 The greenhouse effects and greenhouse gases

The greenhouse effect, despite the negative connotations associated to the concept, is a naturally occurring phenomenon that has enabled life on Earth for millennia.

Sunlight enters the atmosphere in the form of waves. Short-wavelength radiation is partly absorbed by the ozone layer (mostly ultraviolet → about 25%). The remaining 75% of larger wavelengths radiation reaches the Earth's surface, where it is absorbed and produces heat.

The Earth's surface re-emits radiation (like a reflection) at much longer wavelengths (infrared → heat).

A high percentage (75–80%) of this radiation is absorbed by greenhouse gases in the atmosphere.

These gases re-emit radiation, and some of it reaches the surface of the Earth again, causing warming (atmospheric temperature is significantly warmer than outer space).

Greenhouse gases are special in that they have the ability to absorb this long-wavelength radiation that is emitted from the Earth's surface.

The main greenhouse gases are carbon dioxide and water vapour both found in low concentrations in the atmosphere.

Oxides of nitrogen and methane are also greenhouse gases present in the atmosphere.

It is important to realize that ozone is a greenhouse gas, so ozone depletion does not increase the greenhouse effect.

4.4.2 The enhanced greenhouse effect and its effect on global temperatures

Currently, increased levels of greenhouse gases in the atmosphere caused by industrialization cause the atmosphere to retain more and more heat. This is known as the enhanced greenhouse effect, a phenomenon that has affected average global temperatures and climate patterns on Earth.

Human activity has considerably increased the production of greenhouse gases (predominantly carbon dioxide) in the last 200 years, starting after the onset of the industrial revolution, where the combustion of fossil fuels became a major source of fuels and energy for human development.

Some human activity that has led to the increase in greenhouse gas production:

- Burning of fossil fuels
- Use of ammonia-based fertilizers
- Industrial processes (e.g., production of nitric acid)
- Waste disposal in landfills
- Production and distribution of natural gas

These processes have led to a considerable increase in carbon dioxide and methane in the atmosphere. This in turn has caused a major increase in global temperatures in the last two centuries, causing rises in sea level, destruction of Arctic habitats, coral reefs, and much more.

4.4.3 Changes in atmospheric CO₂ concentrations and average global temperature

Over the last 150 years, carbon dioxide emissions have increased significantly, mostly as a result of fossil fuel combustion in industry. While there is a clear correlation between an increase in carbon dioxide emissions and an increase in global temperatures, other variable factors influence this process (e.g., atmospheric concentration of other greenhouse gases).

EVOLUTION AND BIODIVERSITY

5.1. Evidence of evolution and natural selection

– Evidence of evolution – Natural selection

106

5.2. Classification of biodiversity

– Binomial system and dichotomous key – Domains and phylogeny – Classification of plants – Classification of animals – Classifications of vertebrates

110

5.3. Cladistics

– Human classification through cladistics – Reclassification of figworts – Cladogram analysis

116

5.1 Evidence of evolution and natural selection



Evolution is the accumulation of changes in heritable characteristics of a species over time.

This means that over times, certain characteristics of a species change, and if these characteristics are inheritable, then this change is transferred to all the subsequent generations.



Speciation is a process through which one species splits into two separate species.

Adaptation is the process of change by which an organism or species becomes better suited to its environment.

If the members of one species become separated for a long period of time, their characteristics (due to adaptations to the new environment) will start changing as well.

Sometimes it happens that these two populations change so much that, even if they got reunited, they would not be able to interbreed anymore.

5.1.1 Evidence of evolution

Pentadactyl limb

Mammals, birds, reptiles and amphibians all have limbs which they use to different purposes.

Some limbs are used for running, walking, swimming or jumping, but eventually, all possess the same pentadactyl structure (5 digits).

The explanation for this is that all these animals have the same ancestor that had a pentadactyl limb, but as the organisms adapted to different environments (water, forest, desert, etc.), the exact structure started differing.

As the limb structure changed as a response to different environments, this type of evolution is called adaptive radiation.

Selective breeding

Domesticated animals are an example of a “fast forwarded” evolution.

Wild animals with favourable characteristics for humans were bred with other such animals of the same species to get offspring with similar characteristics.

This process is called selective breeding.

An example of this would be a dog, which is a domesticated wolf.

Wolves that were less aggressive were domesticated and bred until more such offspring were produced.

Then, for example, only the smaller individuals were bred which resulted in a smaller dog species (etc.)

Fossil records

Fossils are remaining of organisms found in stones that can help us determine their age and compare them to the currently living species.

It is possible to determine the age of stones where fossils were found and this has shown that bacteria are found in the oldest fossils, followed by algae, fungi and more complex organisms later.

Acanthostega is a fossil of a vertebrae that doesn't match any of the current living species, but shows similarities with them.

It has both 4 limbs that matches amphibians, but also a fish-like tail and gills which suggests it was probably a transition species between the fish and amphibians.

Melanism

Melanism refers to the phenomenon where a lightly coloured species has a darker variant.

Lightly coloured moths are well adapted to avoid predators by blending with the lightly coloured tree branches.

The melanistic form cannot blend in with the trees so are frequently eaten.

During industrial revolution, the smog caused the trees to darken so the lightly coloured moths suddenly became very visible on the tree branches and got eaten more often.

In these areas, the melanistic form was less noticeable so it had higher rate of survival, and became more prevalent.

The switch from the light population to the dark population is an example of evolution as it shows how the species changes as a result of a changing environment by passing on its favourable genes.

5.1.2 Natural selection

Charles Darwin was the first scientist to publish the theory of natural selection, although Alfred Wallace had the same idea at the same time (but was just too slow to publish).

Natural selection suggests that the better adapted species will have a higher chance of survival and will therefore be able to pass on their genes and the species will evolve towards the better adapted species.

Observations of natural selection

There are more organisms than the environment can support

Individuals must compete for natural resources to survive. Different individuals of the same population have slightly different traits which make them better or less well adapted to the environment. Individuals with better adapted traits are more likely to catch prey, survive and breed, so their genes are passed on to the next generation. In this way, the favourable characteristics are increased within the population, and the *population gradually evolves*.

* Note that only the genetically determined traits can be passed on this way. An athlete, whose muscles have grown through exercise will not pass her big muscles to her offspring.

Examples of natural selection

Example.

Beak size of finches

Galapagos finches feed on seeds that fall on the ground, and for that, they have specially adapted beaks.

Some finches have larger, and some finches have smaller beaks. The larger beaks are better for eating larger, harder seeds.

During drought season, smaller seeds are not common, and only the large seeds are produced. If the drought extends for a long period of time, as it did between 1974 and 1977, the finches with larger beaks tend to be better adapted to cracking bigger seeds and are more likely to survive.

The mean beak size during those drought years was found have increased due to the fact that the animals with smaller beaks couldn't open the larger seeds and would therefore die before passing on their genes.

However, during a rainy season in 1983, there were more smaller seeds, and the animals with smaller beaks were faster to pick those up, so they had an advantage compared to the ones with larger beaks.

The mean beak size again decreased, as now the better adapted individuals were those with the smaller beaks, and they could breed and create more offspring.

Example.

Antibiotic resistance in bacteria

Bacterial antibiotic resistance is a growing problem in the world.

Normally, the antibiotic resistance gene exists in organisms that naturally produce the antibiotic.

This gene can be transferred to bacteria through the means of a plasmid.

The bacteria that have the gene cannot be killed by the antibiotic, so when it is administered to the patient suffering from a bacterial infection, all the non-resistant bacteria will die, and the resistant ones will survive and multiply.

Since these bacteria can't be killed by this type of an antibiotic, another antibiotic can be administered, killing this population, but giving a selective advantage to the ones possessing resistance gene for the other antibiotic.

In the end, only the resistant bacteria survive, multiply and exchange genes, leading to a broad range of ineffective antibiotics.

This can be avoided by limiting the antibiotic use, and by always taking the full dose that ensures that all the non-resistant bacteria are killed and that they can't acquire resistance genes from the resistant bacteria.

5.2 Classification of biodiversity

5.2.1 Binomial system and dichotomous key



Binomial system is a universal system used to name the newly discovered species.

The first name in the system is *genus* and is written in the capital letter. The second name is the name of the species, written in the lower-case letter. When printed, the name is written in italics, such as *Homo sapiens*.



Dichotomous key is a tool that helps biologists distinguish between different organisms and classify them correctly.

An example of dichotomous key will be made after you learn about the classifications of plants and animals.

The key makes use of a series of statements about the features of an organism. These statements are numbered. Some statements lead to other statements that help better classify the organism. Other statements immediately give the name of the organism in question



Classification is the term for grouping of species.

Taxonomists are biologists that specialize in the field of classification.

Artificial classification refers to classifying organisms based on one feature that they share, but disregarding all the other ways in which they differ.

An example of this would be classifying all animals with a tail together. A squirrel and a lizard would both be classified together, even though they share very little in common, besides that, and their tails have different structure and function.



Natural classification is an alternative way of grouping organisms, based on their ancestry.

If species are grouped based on their common ancestor, they will share many more features. This way of classification also helps identification of species, and is useful for prediction of their characteristics.

5.2.2 Domains and phylogeny

All organisms are classified into three main domains, namely *archaea*, *eubacteria* and *eukaryota*.

Previously, the organisms were divided either into prokaryotes or eukaryotes, but archaea and bacteria are as different from each other as they are from the eukaryotes.

Based on their RNA sequences, it seems that eubacteria and archaea diverged very early in the evolution.

Viruses are not living organisms so they do not fit into any of the domains.



Taxon is a group of organisms, grouped by the means of natural classification.

The 8 main taxa with the animal and plant examples are:

Table 5.1: 8 main taxa.

Name	Human	Sequoia
Domain	Eukaryota	Eukaryota
Kingdom	Animalia	Plantae
Phylum	Chordata	Coniferophyta
Class	Mammalia	Pinopsida
Order	Primate	Pinales
Family	Hominidae	Cupressaceae
Genus	<i>Homo</i>	<i>Sequoia</i>
Species	<i>sapiens</i>	<i>sempervirens</i>

5.2.3 Classification of plants

Plants are classified into 4 main phyla, and you need to know how to distinguish between them, based on their structure.

Bryophytes – Mosses

- No real roots, but only rhizoids (root-like hairs)
- Simple leaves and stems
- No vascular tissues
- Reproduce using spores which are stored in capsules at the end of a stalk



Filicinophytes – Ferns

- Real roots and leaves, but short non-woody stems
- Leaves are often divided into pairs of leaflets
- Have vascular tissue
- Reproduce through spores which are made in sporangia, inside the leaves



Coniferophytes – Conifers

- Have real roots, leaves and woody stems
- Thick leaves with a waxy cuticle (imagine what a Christmas tree would look like)
- Have vascular tissue
- Reproduce using seeds which are made by female cones in structures called ovules
- Male cones will produce pollen used for fertilization



Angiospermophytes – Flowering plants

- Real roots leaves and stems
- Some plants, like trees and shrubs have a woody stem, others don't
- Have vascular tissue
- Reproduce using flowers - seeds are again made in ovules, but now in ovaries of flowers
- Fruits, which develop from these ovaries are used to disperse the seeds



5.2.4 Classification of animals

We will discuss 7 of the 30 possible animal phyla.

Porifera – Sponges

- No symmetry
- Attached to the surface
- No mouth or anus
- Contain pores



Cnidaria – Jellyfish

- Radial symmetry
- Tentacles and stinging cells
- Mouth, no anus



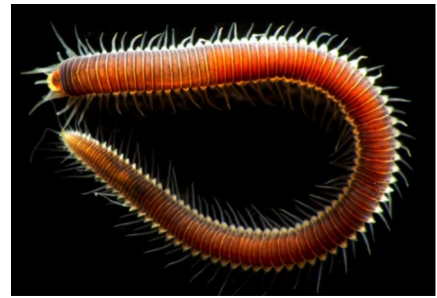
Platyhelminths – Flatworm

- Bilateral symmetry
- Flat body
- Unsegmented
- Mouth, no anus



Annelida – Centipedes

- Bilateral symmetry
- Segmented
- Mouth and anus



Arthropoda – Spiders and scorpions

- Bilateral symmetry
- Exoskeleton
- Segmented
- Appendages divided by joints



Mollusca – Snails

- Mostly bilaterally symmetrical
- Muscular foot and mantle
- Shell
- Mouth and anus



Chordata – Fish, mammals etc.

- Notochord and dorsal nerve chord (imagine a spine and nerves in it)
- Post-anal tail



5.2.5 Classifications of vertebrates

The phylum of chordata are very diverse, and we will now classify them further.

Bony ray-finned fish

Unlike in a dolphin with soft fins, these fish have fins supported by bones.

- Scales on skin
- Gills with one slit
- Fins supported by rays
- Swim bladder (so that they can change altitude in water)
- External fertilization

Amphibians (frogs)

- Soft, permeable skin
- Lungs with few folds
- External fertilization with gel that protects the eggs
- Larva live in water

Reptiles (crocodiles)

- Dry and scaly skin – impermeable
- Lungs with a lot of folds
- Internal fertilization with soft shelled eggs
- One type of teeth

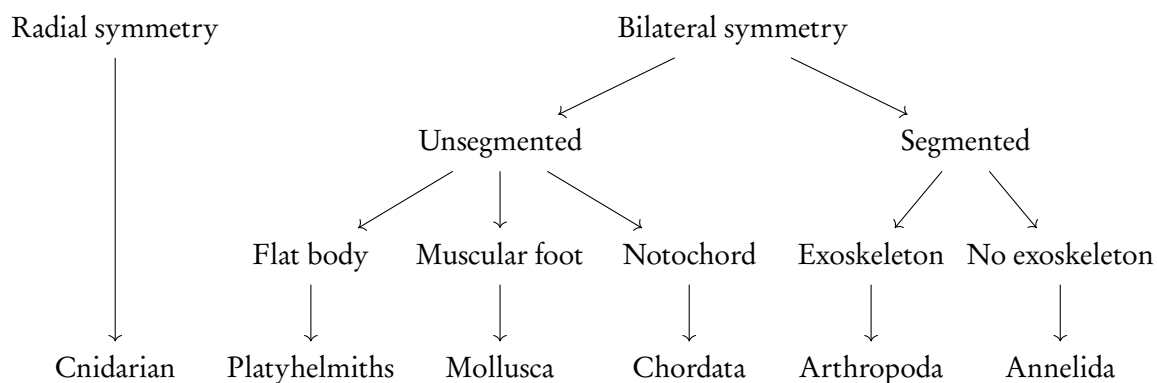
Birds

- Feathers on skin
- Lungs with parabronchi
- Wings, but no front legs
- Hard shells on eggs
- Beak, and no teeth

Mammals

- Hairs on skin
- Lungs with alveoli
- Give birth to live animals
- Mammary glands with milk
- Teeth of different kinds

Let's construct that dichotomous key, based on animal phyla!



5.3 Cladistics



Clade is a group of organisms that evolved from a common ancestor.

Cladogram is a tree diagram that shows how clades diverged one from another.

Cladistics is a method of classification that analyses base and amino acid sequence data to determine ancestry and construct cladograms.

Analogous traits are those traits that have a similar appearance and function, but have do not share the same origin.

Analogous traits arise from convergent evolution, meaning that species of different origins, developed similar features in response to their environment.

An example of this could be the wings of birds and bats, which have the same function even though the species developed these traits independently.



Homologous traits are traits that are structurally different, but have the same origin.

Homologous traits are a result of divergent evolution, where a particular trait, shared by many species, accumulates structural differences as a response to the environment.

An example of this is the previously mentioned pentadactyl limb (first section of this chapter).

In the past, the only way to compare ancestry of certain species was to look at their anatomical features, but at some point, as in the case of bats and birds, it might be difficult to determine whether these features share a common ancestor, or have arisen by convergent evolution.

Today, cladistics takes advantage of possibility of genome sequencing. There is a positive correlation between the number of amino acid sequences, and the time since they have split from a common ancestor.

5.3.1 Human classification through cladistics

Mitochondrial DNA was used to classify humans and several other primates, using a cladogram. Mitochondria contain their own DNA, which is passed on from a mother to her child, as it is the mother gamete's cytoplasm that is used for the first cell of the embryo development. The size of these DNA molecules is much smaller than that of nuclear DNA, and it resembles that of plasmids in prokaryotes (remember the symbiotic theory).

Still there are base pair differences that can help us determine the time points when certain primate species diverged from each other

- 5 million years ago: Human-Chimpanzee
- 140,000 years ago: African-European/Japanese
- 70,000 years ago: European-Japanese

5.3.2 Reclassification of figworts

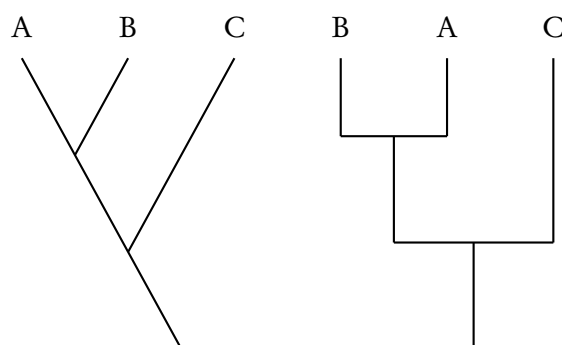
Figworts were originally classified based on their structural differences. Sequencing of their genome showed that species within that family did not share a common ancestor.

Some plants were moved to pre-existing families (like plantain and broomrape). Others were transferred to completely new families based on evidence of a split between species. Some pre-existing families were found to actually share the ancestor with figworts, so these families were all merged into one.

With the developments in computers' analytical power and sequencing methods, there will probably be more and more re-classifications.

5.3.3 Cladogram analysis

Cladograms are constructed by plotting lines with branching points which represent points of divergence between species. These lines are usually plotted against time and percentage difference in the analysed variable (amino acid sequence, base pair sequence etc.).



HUMAN PHYSIOLOGY

6.1. Digestion and absorption 121

– Digestion: why is it essential? – Mixing and moving food along the gut: longitudinal and circular muscle contraction in the small intestine – Digestive enzymes – Absorption: the structure and function of villi

6.2. The blood system 124

– The cardiac cycle: blood circulation in the body – Control of heartbeat: the role of the pacemaker cells, brain signals and adrenalin

6.3. Defence against infectious diseases 130

– Blood clotting – Antibody production – HIV – Human Immunodeficiency Virus

6.4. Gas exchange 136

– Ventilation: why do we need it? – Alveoli: the site of gas exchange – Inhalation and exhalation: muscle contractions needed to regulate airflow

6.5. Neurons and synapses 140

– Action potential: generating and transmitting electrical signals in the brain – Neurotransmitters: chemical signalling across synapses

6.6. Hormones and homeostasis **143**

- Blood glucose – Diabetes – Thyroxine and metabolism
- Leptin and melatonin

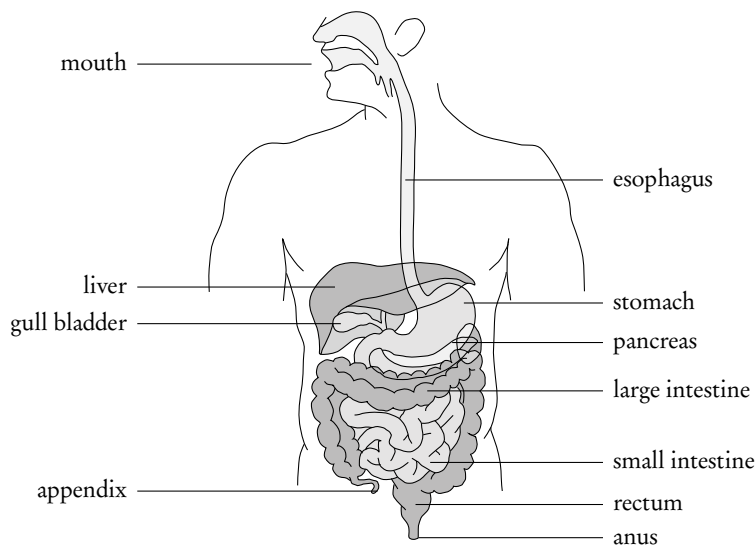
6.7. Reproduction **145**

- Embryo development – Steroid hormones – The menstrual cycle – In vitro fertilization – Harvey's reproduction experiments

6.1 Digestion and absorption

6.1.1 Digestion: why is it essential?

The food we eat consists of large and complex organic molecules, eg. *starch*, which *cannot be absorbed* by cells in the intestines. Digestion is the biochemical *breakdown of large, insoluble molecules into smaller ones*. For example, starch broken down into glucose is a useful source of energy.



Skill: Production of an annotated diagram of the digestive system (TIP: annotated suggests that the elements shown on the diagram must be briefly described, use the information on this section to correctly annotate the diagram of Figure 6.1 with the basic function of each labelled structure)

Figure 6.1: Digestive system.



Chemical Digestion is the breaking down of food with chemical agents. This includes enzymes, acids, and bile.

Mechanical Digestion is the act of physically digesting food through chewing, churning, and segmentation

6.1.2 Mixing and moving food along the gut: longitudinal and circular muscle contraction in the small intestine

Digestion begins in the mouth, where chewing breaks down the food into smaller pieces, as well as some enzymes begin to break down some carbohydrates. Then, the partially digested food arrives to the stomach, where churning occurs, which are movements of the stomach that mix and breakdown molecules more. Here, there are acids that also allow enzymes to get activated and break down molecules, as well as destroy possible pathogens with the low pH. Partially digested food from the stomach enters the small intestine and moves down this structure due to *peristaltic muscle contraction*.

The small intestine has an inner layer of *longitudinal muscle* and an outer layer of *circular muscle*. Circular muscles contract behind the food to prevent backflow, whereas the longitudinal muscles contract to move the food along the intestine. When both layers of muscle contract simultaneously, they allow for the food to be mixed with digestive juices from the gall bladder and pancreas.

To enhance the absorption process, the surface of intestinal cells contain small *finger-like projections* called *villi*, each of which contains a network of capillaries and a *lacteal* (a branch of the lymphatic system that enables lipid absorption) that connect to larger blood vessels and the lymphatic system. Moreover, each cell that lines the intestinal wall has *hair-like extension* called *micro-villi* that further increase surface area for absorption.

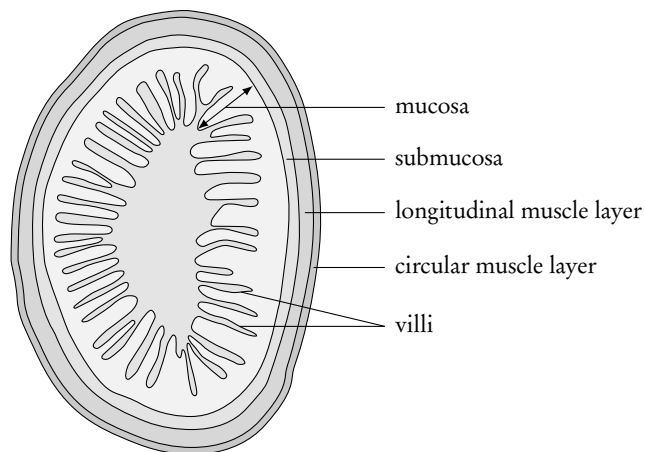


Figure 6.2: The structure of the small intestine.

6.1.3 Digestive enzymes

Physical breakdown of large food molecules is not enough for molecules to be small enough for absorption. Therefore, complementary chemical breakdown is necessary. Enzymes act as *biological catalysts* increasing the rate of digestion. This allows digestion to occur at *normal body temperature*. Several enzymes are needed as each is *substrate specific*. Note the source, substrate, products and optimal pH of each enzyme.

Skill: identification of tissue layers in transverse sections of the small intestine viewed with a microscope/micrograph

Enzyme type	Example	Source	Substrate	Products	Optimal pH
Amylase	Salivary amylase	Salivary glands	Starch	Maltose	7
Amylase	Alpha amylase	Pancreas	Starch	Maltose	7
Maltase	Intestinal maltase	Intestinal wall	Maltose	Glucose	7
Protease	Pepsin	Stomach wall	Proteins	Small polypeptides / amino acids	2–3
Endopeptidase	Trypsin	Pancreas	Proteins	Small polypeptides	7
Lipase	Pancreatic lipase	Pancreas	Triglycerides	Fatty acids + glycerol	7

6.1.4 Absorption: the structure and function of villi

The absorption phase is the process by which the products of digestion, mineral ions and vitamins are taken up through the villi that line the small intestine. To achieve maximal absorption the structure of the villus has a number of functional adaptations:

Villi have a large surface area to volume ration. They are one-cell thick structures, which is advantageous for the products of digestion to easily cross from the lumen of the intestine to the network of capillaries and lacteals for quick absorption.

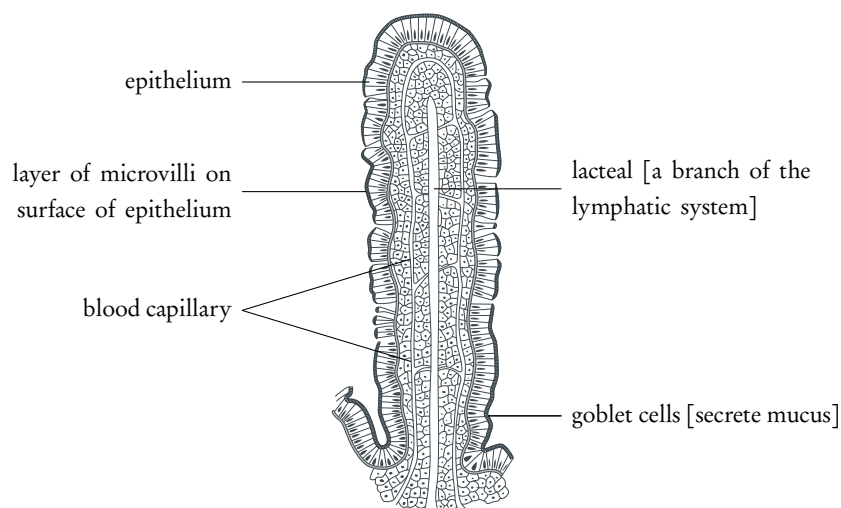


Figure 6.3: Structure of a villus.

Each cell in the mucosa contains smaller structures called microvilli that further increase surface area for absorption of nutrients. These structures contain specific protein pumps and channels that facilitate the movement of molecules across membranes in the correct direction (from intestine to capillaries/lacteals).

Absorption occurs via different processes, all of which have been discussed in previous chapters, depending on the type of molecule to be absorbed:

Facilitated diffusion	(e.g., hydrophilic nutrients like fructose)
Simple diffusion	(e.g., hydrophobic nutrients like fatty acids)
Endocytosis	(e.g., larger molecules like cholesterol and triglycerides)
Active transport	(e.g., charged ions like calcium and sodium)

Example.

Digestion and absorption of starch derivatives

Starch consists of two different molecules: amylose and amylopectin, both linked by alpha-glucose 1,4 links. The only difference is that amylopectin also contains a few alpha-glucose 1,6 bonds. Amylase (salivary and pancreatic) can only break 1,4 bonds, digesting starch into maltose molecules and 1,6 bond-containing segments known as dextrans (which cannot be broken down by amylase). To further digest these molecules, maltase and dextrinase in the small intestine convert the remaining molecules into glucose, which can then be absorbed by the villus through protein pumps (active transport).

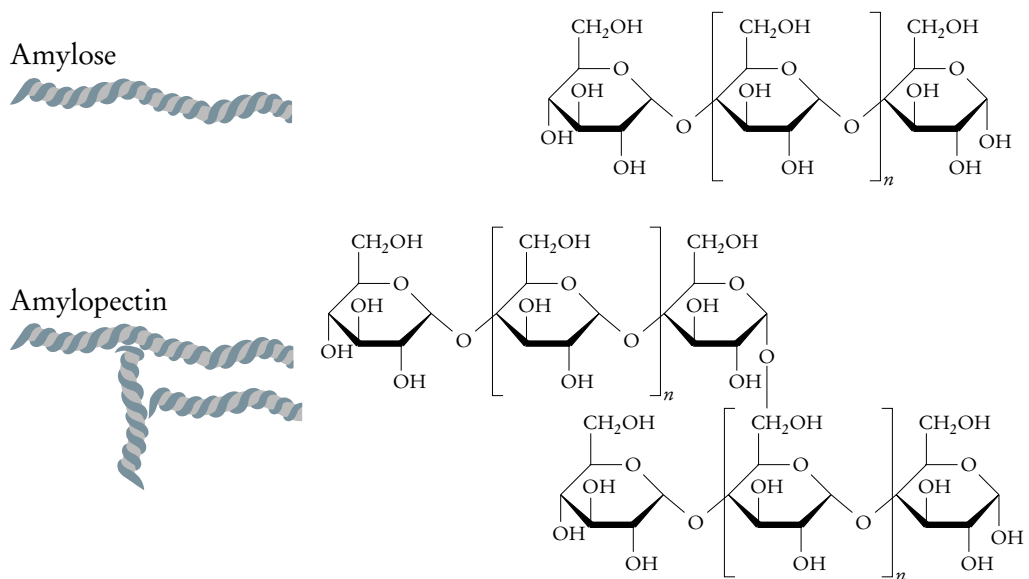


Figure 6.4: Amylose vs. amylopectin.

Diagram showing the difference between amylose and amylopectin (two components of starch molecules). Amylose consists only of 1,4 bonds that result in a straight chain of glucose molecules. Amylopectin on the other hand contains “kinks” that result from 1,6 bonds between certain glucose molecules. These kinks cannot be broken down by amylase, and must be digested by dextrinase in the small intestine.

6.2 The blood system

6.2.1 The cardiac cycle: blood circulation in the body

In the 17th century, William Harvey proposed the theory of blood circulation that continues to be applied today. He demonstrated that the 4-chambered heart was the central “pumping mechanism” that caused blood to circulate the body at high pressures in arteries, and returned to the heart through veins. He also found that these two types of blood vessels are connected by small, hardly visible vessels now known as capillaries. Further research led him to conclude that certain blood vessels contain valves that prevent the backflow of blood, as well as to distinguish between two separate circulations that take place in the body:

Pulmonary circulation that carries deoxygenated blood from the heart to the lungs, where it becomes oxygenated and returns to the heart.

Systemic circulation that carries newly oxygenated blood to the rest of the body, and returns deoxygenated blood back to the heart to enter pulmonary circulation.

Nowadays, the exact mechanism and structures involved in blood circulation have been clearly defined.

Figure 6.5 is a diagram of the heart, our “central pump”, labelled with the major blood vessels, valves, and chambers involved in circulation.

The atria collect blood from veins (vena cava/pulmonary) at low pressure. Blood leaves the atria into ventricles ensuring the ventricles are full.

On the other hand, ventricles pump blood into arteries out of the heart and into the 2 systems previously discussed. They can pump blood at high pressure because of their thicker, muscular walls. The heart valves work with the atria and ventricles to keep blood moving by preventing backflow. Note that the left ventricle supplies blood for the systemic circulation whereas the right ventricle supplies blood for pulmonary circulation.

Skill: Recognition of the chambers and valves of the heart and the blood vessels connected to it in diagrams of the heart structure.

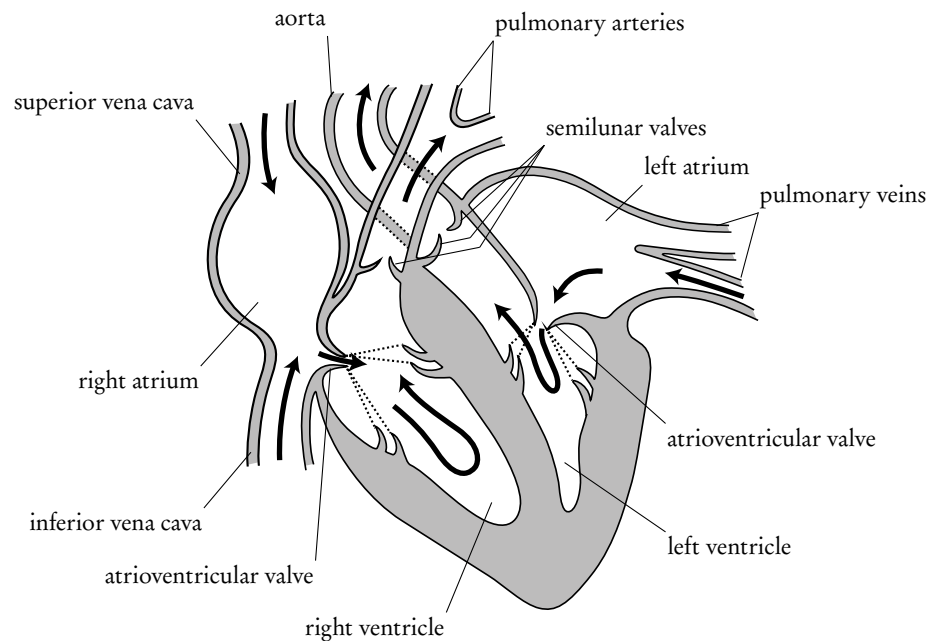


Figure 6.5: Heart structure (cross section).

TIP: Try going through each circulation process using the diagram of the human heart on Figure 6.5!

Pulmonary circulation

- 1 Deoxygenated blood from the superior and inferior vena cava collects within the right atrium.
- 2 The walls of the right atrium contract, pushing blood from the atrium into the right ventricle through the atrioventricular valve.
- 3 Once blood has accumulated in the right ventricle, it contracts powerfully causing:
 1. right atrioventricular valve to close to prevent backflow;
 2. increase of pressure in the right ventricle, leading to the opening of the right semilunar valve (also called pulmonary valve), pumping blood into the pulmonary artery. note that this is an artery that carries deoxygenated blood.

Systemic circulation

Blood from the pulmonary veins collects in the left atrium.

The walls of the left atrium contract, pushing blood from the atrium into the left ventricle through the left atrioventricular valve.

Once blood has accumulated in the left ventricle, it contracts powerfully causing:

1. left atrioventricular valve to close to prevent backflow;
2. increase of pressure in the left ventricle, leading to the opening of the left semilunar valve (also called aortic valve), pumping blood into the aorta.

The left ventricle has a much thicker musculature, as the aorta effectively distributes blood to the entire body, so an even more powerful contraction is necessary

- 4 Blood is carried by arteries, arterioles and capillaries toward the lung alveoli, where it is oxygenated.
- 5 Pulmonary veins return oxygenated blood to the left atrium. Note that these are veins that carry oxygenated blood.
- The aorta branches towards the entire body, one of the first branches directs blood to the coronary arteries (which supply the heart muscle with oxygenated blood for efficient muscle contraction). The rest of the blood is carried by arteries, arterioles and capillaries to the whole body to provide nutrients and oxygen.
- Venules, veins and the inferior and superior vena cava return deoxygenated blood to the right atrium.

Both circulations occur simultaneously, so atrial and ventricular contractions occur in unison. Figure 6.6 is a graph of the pressure changes at each of the processes described previously in the atria and ventricles:

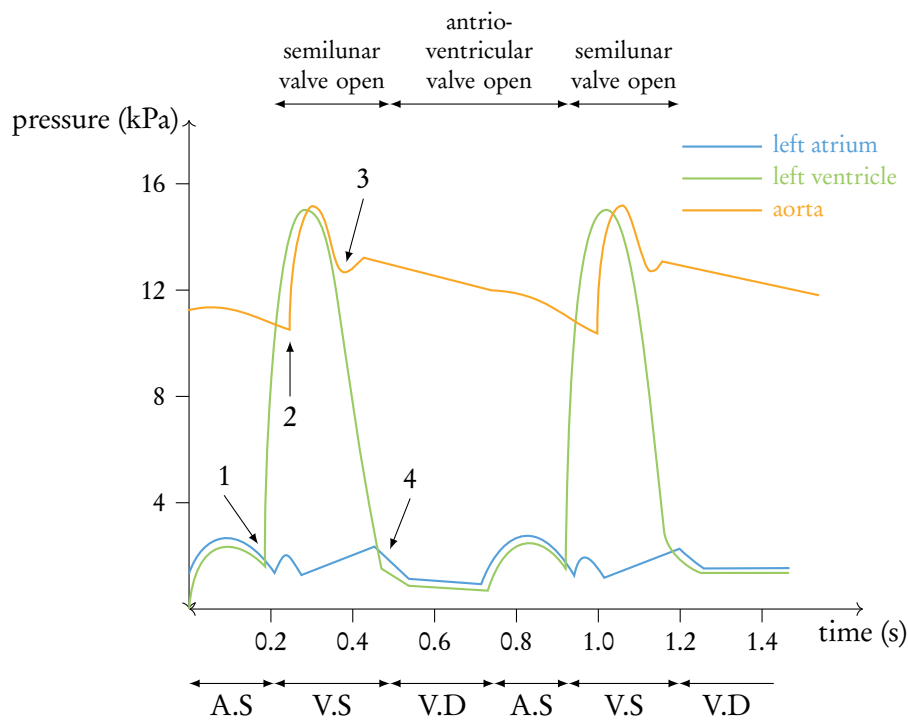


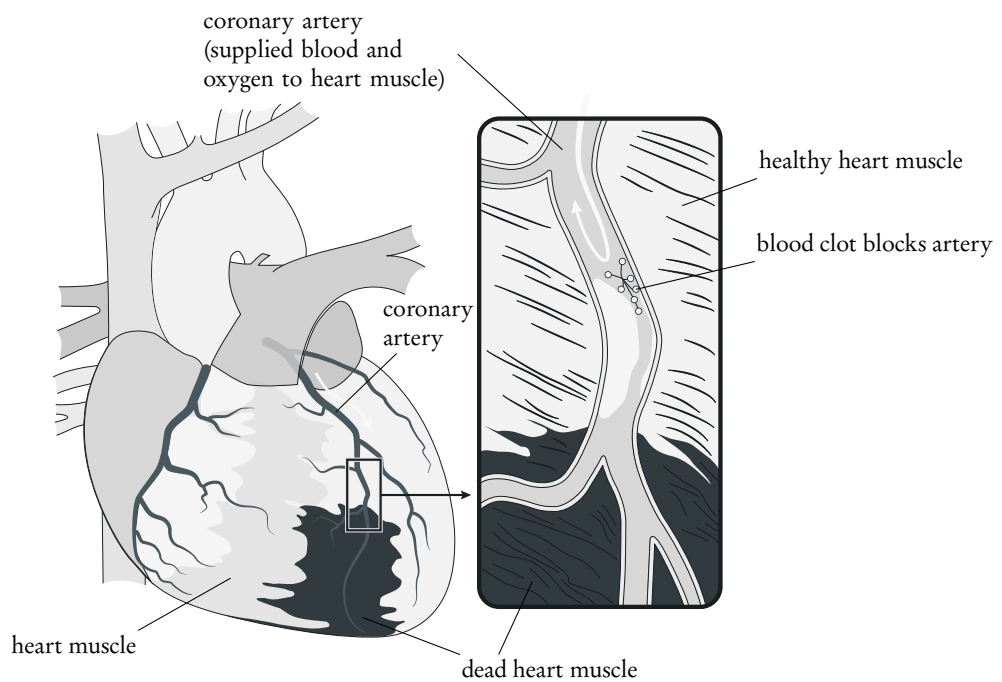
Figure 6.6: Pressure in the cardiac cycle.

Coronary occlusion

Coronary artery occlusion, a common phenomenon caused by fatty plaques building up in the inner coronary arteries is a dangerous occurrence that restricts oxygen and nutrient supply to heart muscle, limiting contraction and thus blood circulation. This can cause chest pain and potential cardiac failure.

Some potential causes for this disease include:

- Hypertension
- Smoking
- High blood glucose (usually due to diabetes)
- High cholesterol levels
- Genetic factors



A comparison of the structure and functions of blood vessels

Artery	Capillary	Vein
Thick walls	Walls one-cell thick	Thin wall
No valves	No valves	Valves
High blood pressure	Low blood pressure	Low blood pressure
Carry blood from the heart	Link arterioles to venules	Carry blood to the heart
Narrow lumen	Narrow lumen	Wide lumen

Skill: Identification of blood vessels as arteries, capillaries or veins from the structure of their walls.

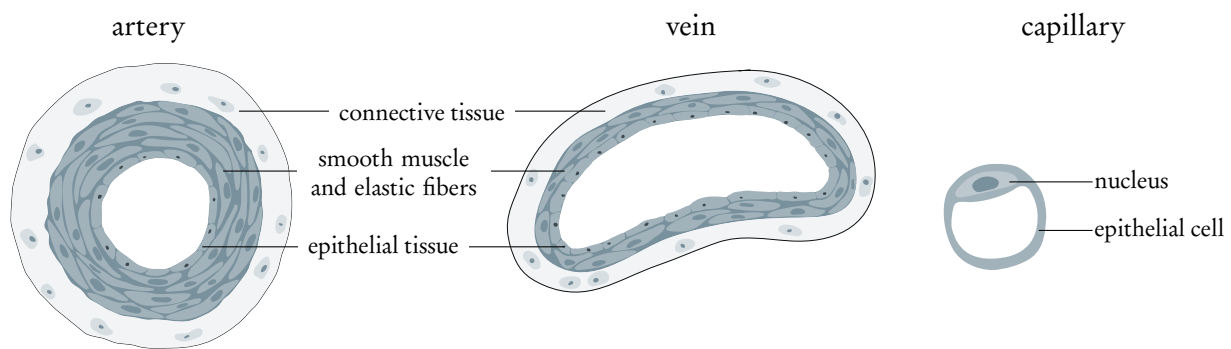


Figure 6.7: Comparison of the structure of an artery, a vein and a capillary.

6.2.2 Control of heartbeat: the role of the pacemaker cells, brain signals and adrenalin

Cardiac muscle is *myogenic*, meaning that it can contract and relax without stimulation from the nervous system. Heart contractions are initiated at an in-built *pacemaker* (which keeps the cardiac muscle working in a coordinated sequence), called the *sinoatrial (SA) node*.

The SA node is a region of pacemaker cells in the right atrium that sets the basic pace of the heart. The SA node produces the initial impulse that *causes both atria to contract at the beginning of each heartbeat*.

This electrical current spreads between cells of the heart to create a coordinated contraction. It first spreads through the walls of the atrium for atrial contraction and gets delayed in between the atria and ventricles at a structure called “bundle of His” to have separate contraction phases. Finally it spreads down through the walls of the ventricles to cause coordinated ventricular contraction.

The *atrioventricular (AV) node* is located at the bottom of the right atrium. This structure also has pacemaker cells, with a rate of firing that is slower than that of the SA node. For this reason, the electrical impulse from the SA node reaches the AV node and sets the pace of the heart beat. If there were to be any dysfunction of the SA node, the AV node could then take over and set a slow “survival” pace of the heart.

The natural rhythm of the pacemaker is modulated by the nervous system (signals from the medulla) and hormones (adrenalin).

Signals to speed up heart rate pass along the sympathetic nerve, and those to slow down heart rate pass through the parasympathetic to slow down heart rate.

Additionally, emotions such as stress and increase in activity level cause the adrenal glands to release the hormone adrenalin (which stimulates the pacemaker to increase heart rate).

6.3 Defence against infectious diseases



Blood clot is a semi-solid lump from liquid blood that is used to seal the cut in the blood vessels and prevent further entry of pathogens into the blood stream.

Platelets are cell fragments present in blood that help create a blood clot upon injury.

Clotting factors are molecules produced by damaged tissues and platelets which set off a cascade of events that lead to the formation of a blood clot.

6.3.1 Blood clotting

In case of a blood vessel injury, platelets and damaged cells release *clotting factors*.

These clotting factors cause the conversion of an inactive protein *prothrombin* into an active form called *thrombin*.

Thrombin further catalyses the conversion of soluble *fibrinogen* to insoluble *fibrin*, which is a long protein that forms a fibrous mesh that catches surrounding blood cells and forms a lump of blood called the blood clot.

If the clot is exposed to air, as is the case at the site of injury, it dries and protects the blood vessels from further entry of pathogens and blood loss.

Blood clotting in coronary arteries

In case of serious plaque deposits in coronary arteries (atherosclerosis), there is a high chance of the plaque rupturing and spilling into the blood stream.

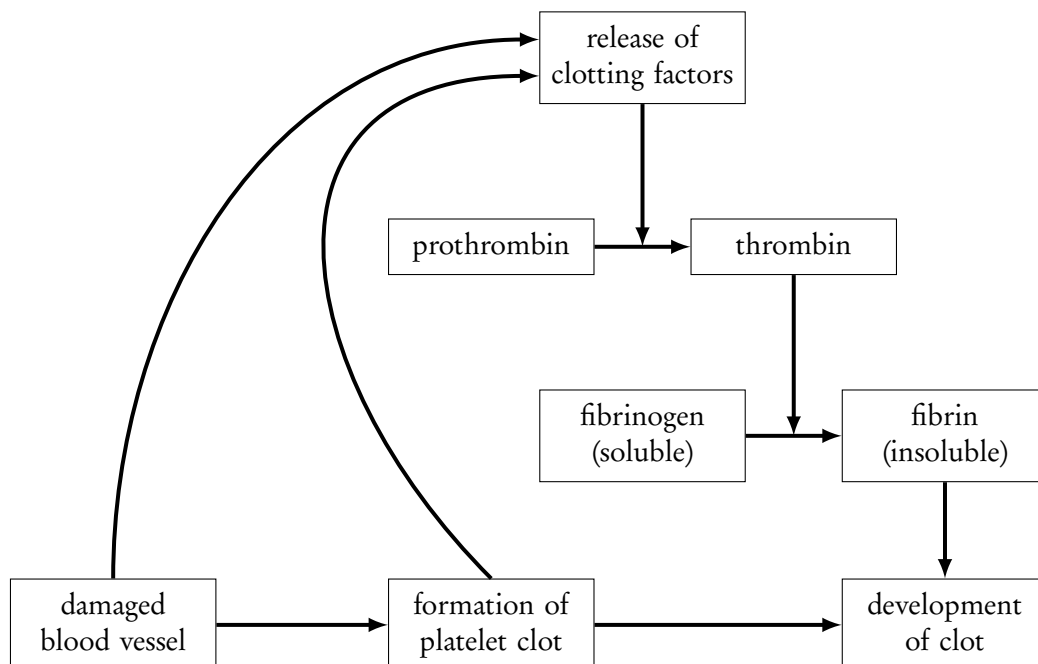
The rupture, as well as the contact of blood with the plaque contents causes a clotting cascade to begin, leading to the formation of a blood clot.

Since coronary arteries are rather narrow, the clot often blocks the blood supply through this artery, and therefore the heart tissue supplied by this vessel stops receiving oxygen and nutrients.

If a blood clot breaks off, it releases a thrombus that can move through the circulation until it gets stuck in smaller arterioles or near capillary beds, cutting off blood supply to the area.

If the supply is blocked for long periods of time the heart tissue gets damaged, which is termed heart attack, or to uncontrolled contractions of the heart called fibrillations.

Some heart attacks are less serious, and the heart can partially recover and start beating again, while more serious artery blocks lead to complete loss of heart function and death.





Pathogen is an organism that causes disease, for example a virus, bacterium or a fungus.

The human body is equipped with two “lines of defence” to protect itself from pathogens:

1st line of defence is the physical barrier, including the skin and mucous membranes.

Besides containing many tough layers as a physical barrier, the skin also provides a chemical barrier in the form of acidic secretions that prevent the growth of pathogens on its surface.

Mucous membranes are parts of the skin covered in a secretion called mucous, that keeps the skin moist and prevents growth of bacteria by killing them with lysozyme enzymes

Mucous membranes can be found in places where the skin has an opening into the body, such as the nose, throat, vagina and urethra.

2nd line of defence is formed by blood cells inside our body.

Phagocytes are white blood cells that ingest pathogens through a process called phagocytosis.

Once ingested, the pathogens are killed by the enzymes in cellular vesicles called lysosomes.

Phagocytes ingest pathogens in the blood and in other tissues, by leaving the capillaries and infiltrating the sites of infections.

Since phagocytes ingest any form of pathogens, they are said to form *non-specific immunity*.

Specific immunity is triggered by other types of white blood cells, called lymphocytes, which produce a response when in contact with a specific type of pathogen

6.3.2 Antibody production



Antigen is any sort of molecule that is recognised by our body as foreign.

Lymphocytes are white blood cells that are involved in specific immunity.

Upon encounter with antigens, lymphocytes can either activate other lymphocytes, or produce antibodies.

Antibody Is a protein produced by lymphocytes in response to the recognition of antigens.

Lymphocytes can only produce one type of antibody, so our body contains many different types of lymphocytes.

Lymphocytes express this type of antibody on their surface so that an antigen can bind it and be recognised by our immune system.

The process of specific immunity goes as follows:

1. When pathogens are encountered, phagocytic cells engulf and break down pathogens. These cells will engulf and break down anything encountered as “non-self”.
2. After engulfment, these “antigen-presenting cells” will display fragments of the broken down pathogen on their surface, and migrate to the lymph nodes to alert the immune system of a threat.
3. Certain lymphocytes called “T-cells” will get activated by this warning and release cytokines to activate particular B lymphocytes.
4. The activated B lymphocytes are those capable of producing the specific antigens for the recognised antigen that was presented by the initial phagocyte.
5. These activated B lymphocytes will rapidly divide into high numbers of short-lived plasma cells that release large numbers of antibodies into the blood stream to target the pathogens for destruction.
6. A small proportion of the activated B and T cells will become memory cells, which provide long lasting immunity to the pathogen. Therefore, in the case of a subsequent infection, these cells will quickly react to produce large numbers of antibodies much faster.

6.3.3 HIV – Human Immunodeficiency Virus

HIV is a virus that reduces the effectiveness of the immune system by reducing the number of active lymphocytes.

HIV does so by penetrating the lymphocytes and integrating their RNA into the cell's DNA with the enzyme reverse transcriptase (which the virus carries)

This viral DNA not only allows the virus to replicate with the cells machinery, but it also leads to a impaired ability to produce antibodies, thus the infected person has a lowered immunity and is more susceptible to infections.

Once the body has lost the majority of this sort of lymphocytes, the condition is termed AIDS, *acquired immunodeficiency syndrome*, since the body lacks the immunity formed by antibody production.

AIDS can culminate in death, as the body cannot fight even the most common infections such as the common cold.

Since HIV is a virus, it cannot survive long outside the body, so it is transmitted through certain bodily fluids:

- Through blood in hypodermic needles (often in drug abusers).
- Through unprotected vaginal, oral or anal sexual intercourse.
- Through the placenta or breast milk.
- Through blood transfusions.



Antibiotics are chemicals produced by certain microorganisms to kill bacteria.

Penicillin is an antibiotic obtained from a *Penicillium* fungus which uses it to kill bacteria that might potentially invade it.

Antibiotics kill prokaryotic organisms (bacteria), and not eukaryotic (plant and animal), because they target metabolic processes specific for prokaryotes.

Therefore, antibiotics can kill bacteria in our body, without harming our own cells.

Viruses cannot be killed by antibiotics because they use the metabolism of the host cell, and therefore, to kill a virus, the cell containing it would also have to be killed.

With different types of antibiotics, most bacterial diseases can be contained, but due to the emergence of antibiotic resistance, some strains of bacteria cannot be so easily killed.

Example.

Florey and Chain

Florey and Chain tested penicillin on 8 mice that were infected with a pneumonia causing bacterium.

The treated mice recovered, while the untreated ones died.

Florey and Chain then treated the same antibiotic on patients dying from bacterial infections, and these patients recovered.

Today, drugs have to go through much more rigorous testing, before they can be given to humans.

First, the safety is thoroughly tested on animals.

Then, healthy humans are given the drug to assess if it is tolerated well.

If all of this goes well, a small number of very sick patients are given the drug, and if this is successful too, only then, a large scale study can be performed.

6.4 Gas exchange

6.4.1 Ventilation: why do we need it?



Ventilation is the exchange of air between the atmosphere and the lungs, achieved by breathing

Gas Exchange is the exchange of oxygen and CO_2 between the bloodstream and the alveoli. This exchange occurs via passive diffusion.

Ventilation is essential because, in order for gas exchange to occur, a concentration gradient must be established (gas with higher O_2 content must come into the lungs in order for it to diffuse into the alveoli, and CO_2 to diffuse out). Gas exchange is passive while ventilation is active to maintain the gradients needed for passive diffusion by continuously refreshing the air in the lungs.

Oxygen is essential for the processes like cell respiration.

Figure 6.8 is a diagram of the respiratory system, air flows through the trachea, bronchi and bronchioles until reaching the alveoli at the tips of the bronchioles, where gas exchange occurs between alveoli and adjacent capillaries.

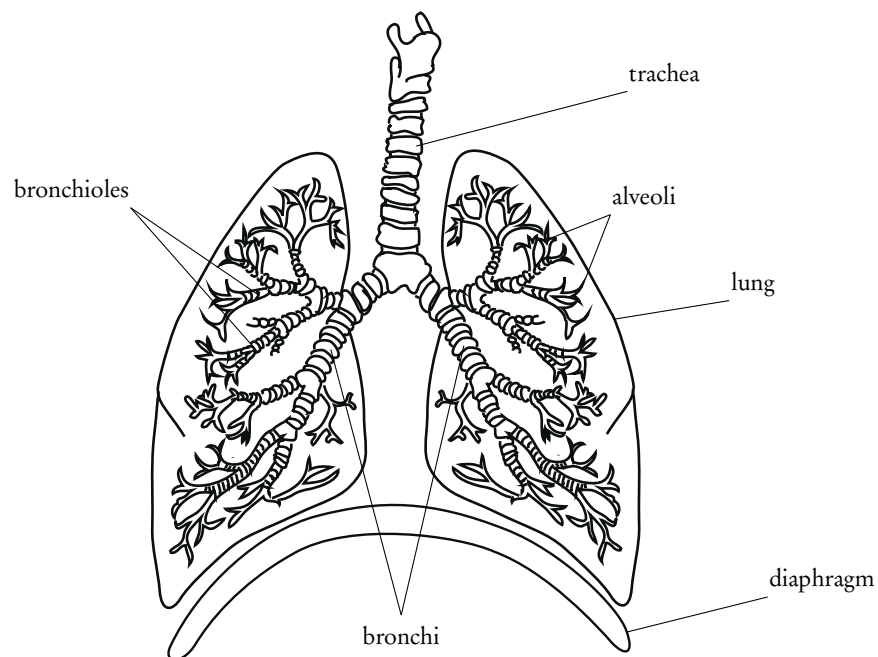


Figure 6.8: Ventilation system.

Example.

Lung cancer: causes and consequences**Causes:**

- Smoking: tobacco smoke contains mutagens that can lead to tumour formation.
- Passive smoking: exhaled smoke from smokers passes carcinogens to others.
- Air pollution: e.g., nitrogen oxides from vehicles, diesel exhaust fuels.
- Radon gas.
- Asbestos and silica: dust from these materials causes cancer if deposited in the lungs.

Consequences:

- Difficulties with breathing.
- Chest pain.
- Persistent coughing.
- Loss of appetite.
- Weight loss.
- Coughing up blood.
- General fatigue.
- Can be fatal.

6.4.2 Alveoli: the site of gas exchange

Alveoli are the body's gas exchange surfaces, formed in clusters at the ends of the smallest bronchioles. They are essentially air sacs with a very small diameter. The presence of many alveoli creates a large total surface area for gas exchange. Each alveolus is surrounded by a network of blood capillaries allowing oxygen to diffuse through into the blood and carbon dioxide to diffuse out in the opposite direction. Here are some additional structural characteristics that enhance this process:

- Two types of cells:

Pneumocytes type I extremely thin and adapted for gas exchange.

Pneumocytes type II secrete a solution containing surfactant, which creates a moist surface inside the alveoli to prevent the sides of the alveolus adhering to each other by reducing surface tension.

- Cells in alveoli are one-cell thick and in close proximity to blood capillaries, which are also one-cell thick. This is essential as it creates a very short distance for diffusion of Oxygen and Carbon Dioxide.

Diagram showing type I and II pneumocytes, as well as the associated capillary. Direction of gas exchange is also shown.

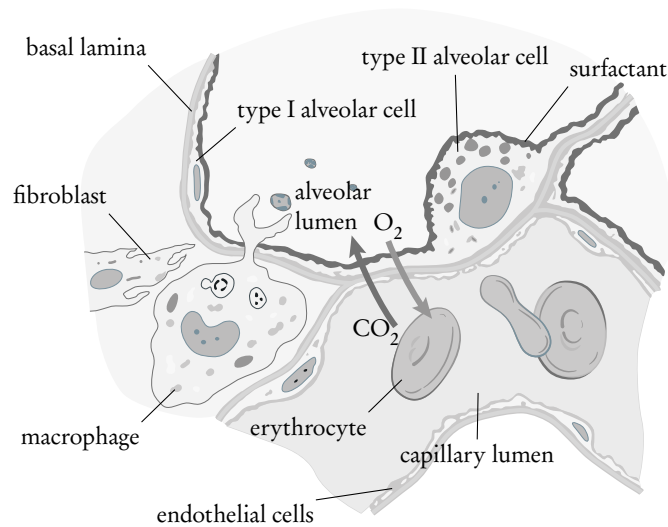


Figure 6.9: Type I and II pneumocytes and capillary.

Example.

Emphysema: causes and consequences

Emphysema is a chronic and progressive respiratory disease. Cilia that line the airways and expel mucus become damaged and cease to function appropriately, leading to a build-up of mucus that can lead to inflammatory response upon inhalation of smoke and air pollution. A protease is released from inflamed cells, causing the digestion of elastic fibres in the lung and eventually, alveolar walls collapse.

Causes:

- Smoking.
- Air pollution.

Consequences:

- Loss of elasticity of the lungs.
- Reduced surface area for gas exchange.
- Difficult to exhale air → difficulty carrying out intense pH.

6.4.3 Inhalation and exhalation: muscle contractions needed to regulate airflow

In order to understand gas exchange, you need to keep in mind that molecules will always flow from an area of high concentration to low concentration by diffusion

During inhalation

The external intercostal muscles contract while the internal intercostal muscles relax thus pulling the rib cage upwards.

The diaphragm contracts and flattens which increases the volume of thoracic cavity.

As a result, there is a reduction in pressure inside the thoracic cavity, leading to air entering the lungs.

During exhalation

In passive exhalation, external intercostal muscles and diaphragm relax, reducing the volume of the thoracic cavity, increasing the pressure of the cavity and therefore releasing air out of the lungs.

Forced exhalation also includes the contraction of internal intercostal muscles, further reducing the thoracic cavity volume.

Table 6.1

	Diaphragm	Internal intercostal	External intercostal	Volume	Pressure
Inhalation	Contracts	Relaxes	Contracts	High	Low
Exhalation	Relaxes	Contracts	Relaxes	Low	High

6.5 Neurons and synapses

6.5.1 Action potential: generating and transmitting electrical signals in the brain

Neurons are the essential messenger units in the body. They transmit electrical impulses from the brain to the rest of the body and vice versa, and transmit information within the brain. Figure 6.10 is a diagram of a myelinated motor neuron, which stimulates muscle fibres to contract.

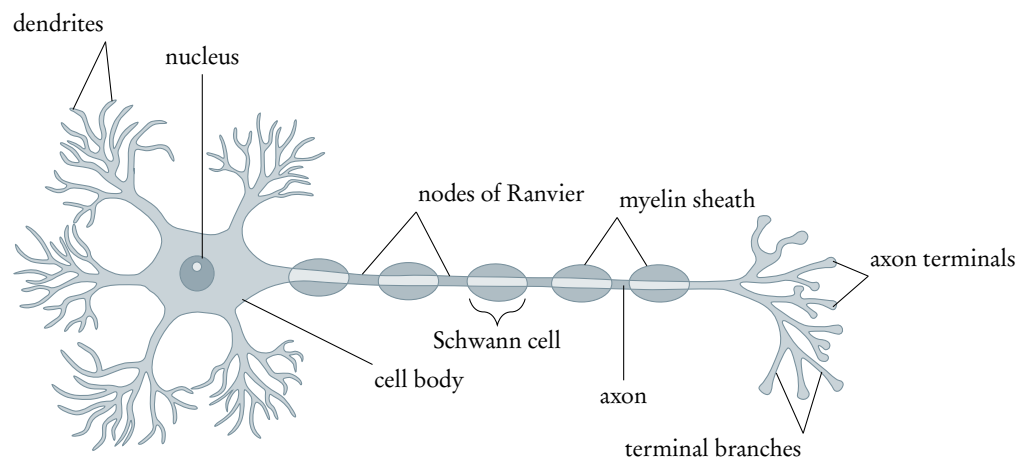


Figure 6.10: Motor neuron structure.

The generation and transmission of electrical impulses across a neuron is achieved by an action potential.



Resting potential The charge difference across the membrane when a neuron is not firing (-70 mV), as maintained by the sodium-potassium pump.

Action potential is an all or none event that consists of rapid changes in charge across the membrane that propagate along an axon when a neuron is firing.

Nerve Impulses are action potentials propagated along the axon of a neuron.

1. Before an action potential can be initiated, the neuron is at resting potential (-70).
2. Resting potential is established the active pumping of K^+ ions into the neuron and of Na^+ ions out of it, creating concentration gradients for both ions, and an overall

negative charge inside the neuron. This is because, as you may remember, the sodium-potassium pump pumps 2 potassium for every 3 sodiums.

3. When input from a previous neuron is stimulating enough and the beginning of the axon reaches its threshold voltage (-55), an action potential is initiated.
4. Depolarization: Sodium voltage-dependent channels open and Na^+ diffuses into the neuron down its concentration gradient, reducing membrane potential and causing more sodium ions to open (membrane potential becomes more positive).
5. Repolarization: Potassium voltage-dependent channels open after a short delay due to the change in membrane potential, and K^+ ions diffuse out of the neuron down its concentration gradient. The inside of the neuron once again becomes more negative. In this stage, the electrical impulse has passed this section of the axon, and the sodium ions in the section adjacent open to transmit the impulse.
6. Concentration gradients of Na^+ and K^+ across the membrane are restored by the Na^+/K^+ pump, and the membrane in this section of the axon once again reaches resting potential (and is once again ready for another action potential).

In myelinated neurons (as seen in Figure 6.10), this ion exchange only occurs at the Nodes of Ranvier, thus conduction of the action potential “jumps” down the axon far more quickly than in unmyelinated axons, where numerous adjacent sections of the axon must undergo the process outlined previously to carry the electrical impulse down the length of the axon. This is known as “saltatory conduction”.

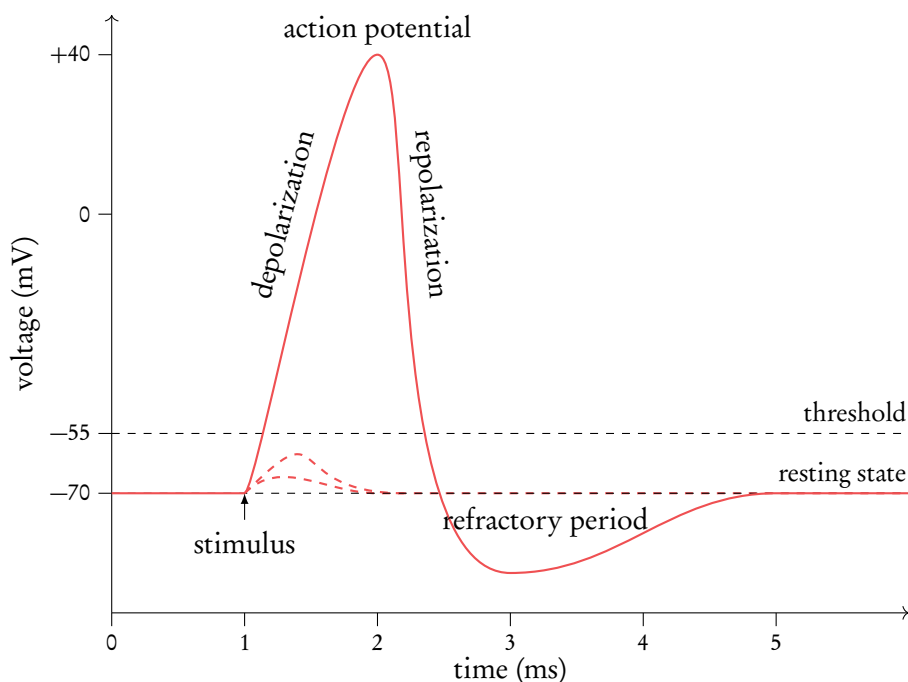


Figure 6.11: Action potential graph showing the processes of depolarization, repolarization and the re-establishing of membrane potential.

6.5.2 Neurotransmitters: chemical signalling across synapses

Once an action potential has reached the axon terminals, chemical signals are necessary to pass on information from the presynaptic neuron to the postsynaptic one or onto an effector cell (e.g., muscle fibre). Separating each neuron are small junctions known as synapses, through which specific chemical messengers called neurotransmitters (NTs) can diffuse. An overview of this process is:

1. A nerve impulse reaches the axon terminal of the presynaptic neuron, leading to depolarization at the terminal.
2. Depolarization leads to the opening of voltage-gated calcium channels and an influx of Ca^{2+} into the presynaptic neuron.
3. Influx of calcium causes neurotransmitter-filled vesicles to fuse with the presynaptic membrane and release NTs into the synaptic cleft by exocytosis.
4. NTs diffuse across the synaptic cleft and bind to specific NT receptors in the postsynaptic neuron.
5. This binding results in the opening of ligand-gated ion channels that allow either Na^+ or Cl^- to diffuse into the postsynaptic neuron.
 - (a) Na^+ influx creates an excitatory signal at the postsynaptic membrane, and allows for the transmission initiation of the action potential if the threshold potential is reached.
 - (b) Cl^- influx on the other hand, leads to hyperpolarization (inside of the neuron becomes more negative), resulting in an inhibitory signal that may prevent action potential initiation.
6. NTs in the synaptic cleft are rapidly degraded by enzymes or are taken up by the presynaptic neuron once more. Ca^{2+} is pumped out of the presynaptic neuron as well to re-establish a concentration gradient.

Example.

Cholinergic synapses

A common example of a synapse is the cholinergic synapse, which most commonly results in the opening of Na^+ ion channels in the postsynaptic membrane and thus transmits an excitatory signal. Acetylcholine is the NT that is released in cholinergic synapses, which then diffuses and binds to nicotinic receptors at the postsynaptic neuron. Enzyme cholinesterase rapidly degrades this NT where one of the residual components, choline, is reabsorbed into the presynaptic neuron.

Some chemicals found in neonicotinoid pesticides have a similar chemical structure as acetylcholine, and can bind to the acetylcholine receptors in cholinergic synapses. Cholinesterase is unable to degrade this chemical and thus, the cholinergic receptor remains blocked and nerve impulses cannot be mediated. This is a very dangerous phenomenon that is lethal for bees and insects that come into contact with these types of pesticides.

6.6 Hormones and homeostasis

6.6.1 Blood glucose

Glucose concentration in blood has to be tightly regulated in order to provide cells across the body with enough energy, but at the same time, to control water movement in and out of the cells.

The pancreas is the organ in charge of glucose regulation, and it does it with the production of two hormones:

Glucagon is produced by *alpha cells* and is secreted when the glucose levels are low:

- it leads to breakdown of glycogen in the liver
- this leads to the release of glucose in the blood

Insulin is produced by *beta cells* and is secreted in response to high blood glucose levels:

- it stimulates cells to take up the glucose from blood;
- it also stimulates liver and muscle cells to store glucose in the form of glycogen.

6.6.2 Diabetes

Diabetes mellitus is a disease of faulty glucose regulation. Two types of diabetes can be differentiated, based on the cause.

Type I diabetes

early-onset diabetes usually develops in those under 20 years old

inability to produce sufficient quantities of insulin

target cells remain sensitive to insulin

is linked with

- genetic predisposition
- virus
- autoimmune disorder
- destruction of (pancreatic) beta cells involved
- requires daily injections of insulin
- beta cell transplant

Type II diabetes

adult-onset diabetes usually occurs in those over 40 years old

inability to respond to insulin mainly due to insufficient receptors on target cells

target cells less sensitive to insulin

is linked with

- dietary
- lifestyle factors
- increased fatty acids in blood
- controlled by diet
- exercise
- weight loss
- medication but not insulin injections

6.6.3 Thyroxine and metabolism



Thyroxin is a hormone produced by the thyroid gland which is responsible for the metabolic activity of the cells.

Thyroxin is made out of 4 molecules of iodine. Low iodine intake can result in lack of thyroxin synthesis (which is why all the salt is additionally iodized).

When the body temperature drops, this is sensed by the hypothalamus in the brain which triggers the production of thyroxin by the thyroid.

Thyroxin increases the metabolic rate of the cells, causing higher heat productions, but also leads to vasoconstriction of skin blood vessels (to preserve heat) and shivering.

6.6.4 Leptin and melatonin



Leptin is a hormone secreted by adipose (fat tissue) cells, which controls appetite.

Leptin is produced by the adipose cells and travels to the hypothalamus in the brain, where it stimulates certain cells to signal for a decrease of appetite.

In this way, the body is able to regulate its body mass, by decreasing food intake as fat tissue increases.



Melatonin is a hormone produced by the pineal gland and it is responsible for controlling the sleep-wake cycles in humans.

In response to different amounts of light/dark, cells in the eye send impulses to the supra chiasmatic nuclei (SCN) in the hypothalamus.

The SCN controls melatonin production, by increasing it at night, and decreasing it at dawn.

Over the years, melatonin production in humans decreases so our sleep cycles become less regular.

When travelling long distances by plane, the SCN and pineal gland cannot adjust to the new light cycles at the new time zone rapidly enough, so during the first few days, the traveller's body continues to function at the old rhythm. This is termed *jet lag*. Taking oral doses of melatonin at the newly designated sleeping schedule can help the body adjust to the new time zone more rapidly.

6.7 Reproduction

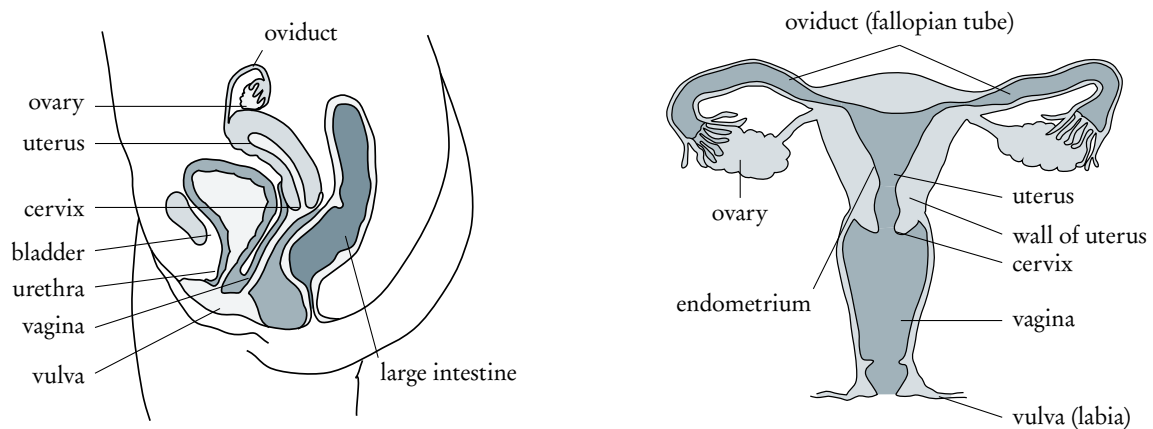


Figure 6.12: Female reproductive system.

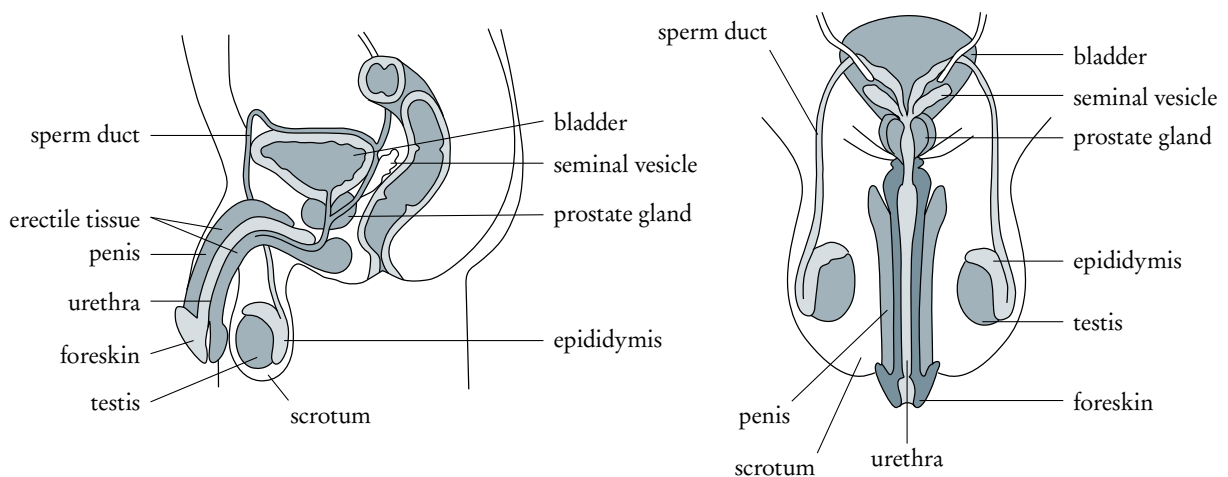


Figure 6.13: Male reproductive system.

6.7.1 Embryo development

The difference between a male and a female zygote is in the 23 pair of chromosomes.

A male has an XY combination, while the female has an XX combination.

A gene called SRY is present in male zygotes, but not in female ones.

If the SRY gene is present, it codes for a testis determining factor protein which causes development of testes.

6.7.2 Steroid hormones

Testosterone is produced by testes, both in a foetus and later in life. In a foetus, testosterone leads to development of male genitalia. During puberty, it leads to development of secondary sex characteristics (growth of pubic hair, penis and testes). In later life, testosterone stimulates sperm production.

Oestrogen is produced in the ovaries. In foetus, it causes development of female genitalia (if there is no testosterone). In puberty, it leads to development of secondary sex characteristics.

Progesterone is also produced in the ovaries and its function is to thicken the uterus before embryo implantation.



Negative feedback means that the increase in a product of a process will lead to the decrease in the process itself.

Positive feedback means that the production of the final product will stimulate the process of production even further.

6.7.3 The menstrual cycle

In order to fully understand the processes during the menstrual cycle, the following terms should be clear.



Pituitary hormones are hormones produced in the pituitary gland (FSH and LH).

Follicle is a growing egg and surrounding fluid and cells

FSH (follicle stimulating hormone) stimulates follicle development and secretion of oestrogen by the follicle.

LH (luteal hormone) causes ovulation, the release of the egg from the ovary. A surge of LH occurs once during the menstrual cycle, and it happens due to a high concentration of estrogen.

Corpus Luteum is the structure left behind after the egg is released from the follicle. It produces progesterone and some estrogen and functions for a limited amount of time after ovulation if there is no fertilization.

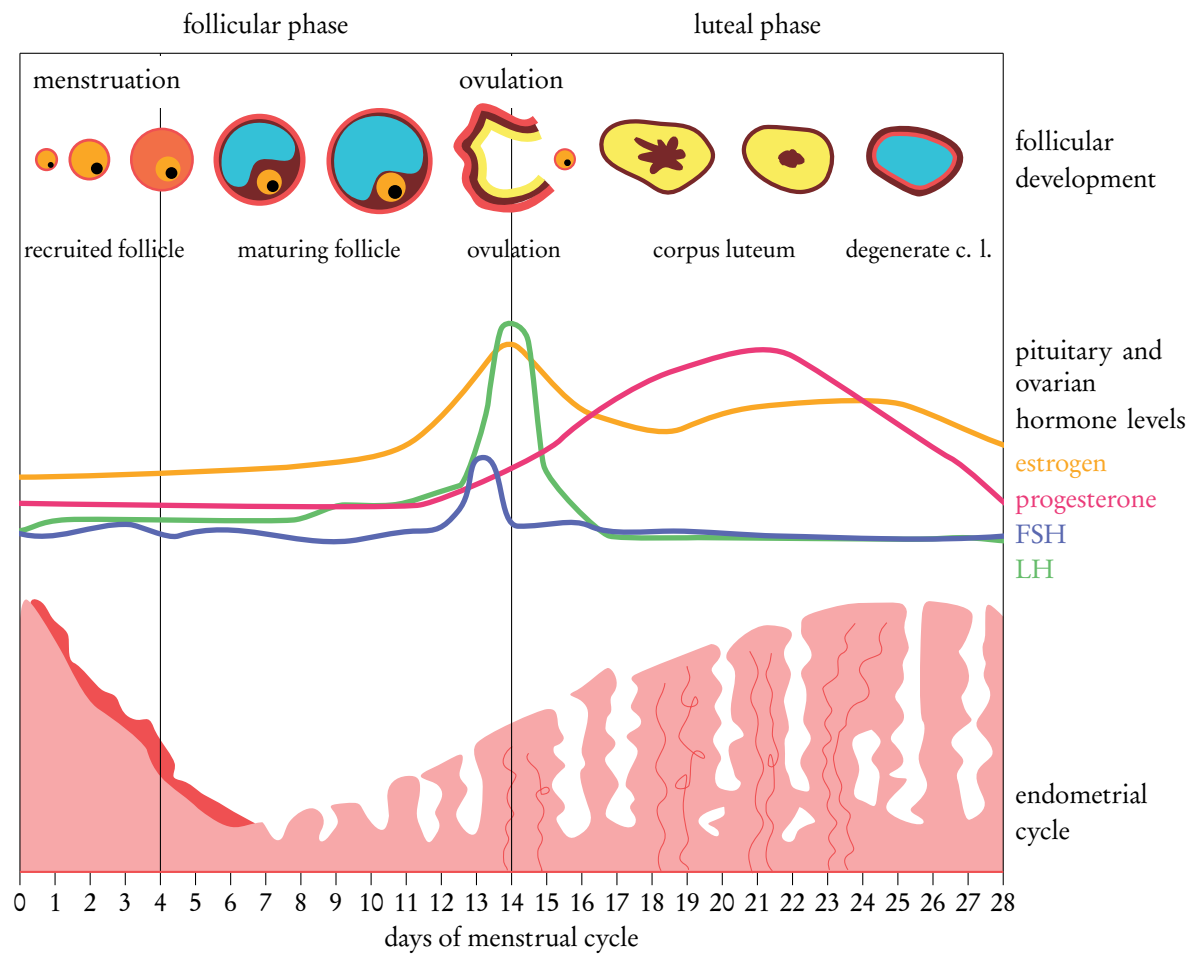
Ovarian hormones are hormones produced in the ovaries (oestrogen and progesterone).

Oestrogen is produced in a positive feedback loop with FSH by the follicle. It causes repairing and thickening of the uterus lining and causes LH secretion. At very high levels, it inhibits FSH, turning into a negative feedback mechanism.

Progesterone is produced by the corpus luteum and maintains the thick lining of the uterus. It inhibits LH and FSH hormones.

The aim of the menstrual cycle is to mature and release an egg, ready for fertilization, and to prepare the uterus for embedding the embryo in its wall.

1. The first, day 0 of the menstrual cycle, is the first day of bleeding. This corresponds to the beginning of the **follicular phase**.
 - (a) FSH increases and stimulates the follicles with the egg to develop.
 - (b) The development of follicles leads to oestrogen secretion.
 - (c) Oestrogen and FSH are in positive feedback because oestrogen production makes follicle cells more sensitive to FSH, which in return results in more oestrogen production.
 - (d) In this way, both oestrogen and FSH increase.
 - (e) Oestrogen leads to repair and thickening of the uterus lining.
2. Around day 14, the uterus has gotten thicker, and the follicular phase of development is ending.
 - (a) When oestrogen is high enough, it causes an LH surge by the pituitary gland. This corresponds to the **luteal phase**.
 - (b) At the same time, the peak of oestrogen lead to FSH inhibition, which in turn leads to a decrease in oestrogen (negative feedback).
 - (c) As LH reaches a peak, it causes completion of meiosis in the egg and release of the egg into the oviduct-ovulation (around day 14).
3. After ovulation, the broken follicle maintains the uterine lining, in case fertilization occurs.
 - (a) LH stimulates the burst follicle to develop into corpus luteum.
 - (b) Corpus luteum produces progesterone, and some oestrogen.
 - (c) Progesterone will continue rising for a few days in order to maintain the thickening of the uterine lining.
 - (d) The increase of progesterone and oestrogen levels inhibits the FSH and LH production by the pituitary (thereby preventing another ovulation).
4. If no fertilization occurs, the corpus luteum starts degenerating, and the menstruation will begin.
 - (a) With degeneration of corpus luteum, progesterone and oestrogen levels start falling, thereby lifting the inhibition of FSH.
 - (b) The drop in progesterone results in degeneration of the uterine lining and therefore, menstrual bleeding.
 - (c) As FSH levels can begin to rise again, a new cycle of follicular development can start.



You should be able to understand the diagram of the menstrual cycle.

6.7.4 In vitro fertilization



In vitro fertilization (IVF) is a fertilization procedure outside of the body, where an egg and sperm cell are combined in a Petri dish to make a zygote. It is a method to overcome issues such as infertility and uterine problems.

The procedure of IVF can be described as the following:

1. Drugs are used to down-regulate the menstrual cycle.
2. FSH is injected to the female to stimulate many follicles to develop.
3. HCG is injected to the female to cause the follicles to mature.
4. The eggs are harvested from the ovaries and a semen sample is collected from the male.
5. The sperm cells are mixed with the mature follicles in a dish outside the body to allow for fertilization.
6. The dish is incubated at 37°C allowing the embryos to develop sufficiently for implantation.
7. The dish examined to choose healthiest embryo.
8. A few healthy embryos are placed in the uterus of the mother using a catheter. Up to four embryos may be implanted, which is why some IVF mothers have more than one child from the procedure.
9. Finally a pregnancy test used to see if procedure has been successful.

Used in cases of

- blocked oviduct;
- low sperm count;
- need for genetic screening;
- infertility;
- cannot become pregnant;
- need for donor embryo.

6.7.5 Harvey's reproduction experiments




William Harvey was a scientist in the 17 century who wanted to explain how copulation resulted in offspring. He chose to study deer during their mating season. He would kill the deer after the mating, in order to observe the development of offspring in the uterus at different stages after mating.

The theory at the time suggested that the male produces an egg that is activated by menstrual blood, and then develops into a foetus in the human uterus.

Harvey showed that this theory was wrong, but he couldn't show that the foetus development was a result of copulation, since the developing embryo was too small to see without a strong microscope.

The story of William Harvey shows how science can only progress with a simultaneous progress in the apparatus and scientific techniques.

NUCLEIC ACIDS

	7.1. DNA structure and replication	154
	<ul style="list-style-type: none">– Nucleosomes: bundles of DNA and protein– DNA replication– Non-coding DNA	
	7.2. Transcription and gene expression	156
	<ul style="list-style-type: none">– Transcription– Post-transcriptional mRNA modifications– Gene expression: regulation of transcription	
	7.3. Translation	158
	<ul style="list-style-type: none">– Translation– Free vs. bound ribosomes	

7.1 DNA structure and replication

7.1.1 Nucleosomes: bundles of DNA and protein

Nucleosomes are the units into which chromatin is organized in the eukaryotic nucleus. One nucleosome consists a globular structure, with a core of 8 histone proteins with DNA wrapped around them. Histone H1 links DNA to the protein core, and each “bundle” of DNA is linked to the next by linker histones. The N-terminal tails of the histones extend outwards from the core of the nucleosome and can attach to neighbouring nucleosomes, pulling them closer together and enabling these to supercoil during the condensation of chromosomes in mitosis. Additional protein modification to the histones (e.g., methylation and acetylation) affects their level of condensation. This serves to influence processes like transcription, by making the DNA available for enzymatic action, or supercoiling to prevent enzyme access. Molecular visualization software can be used to analyse the association of protein and DNA in nucleosomes.

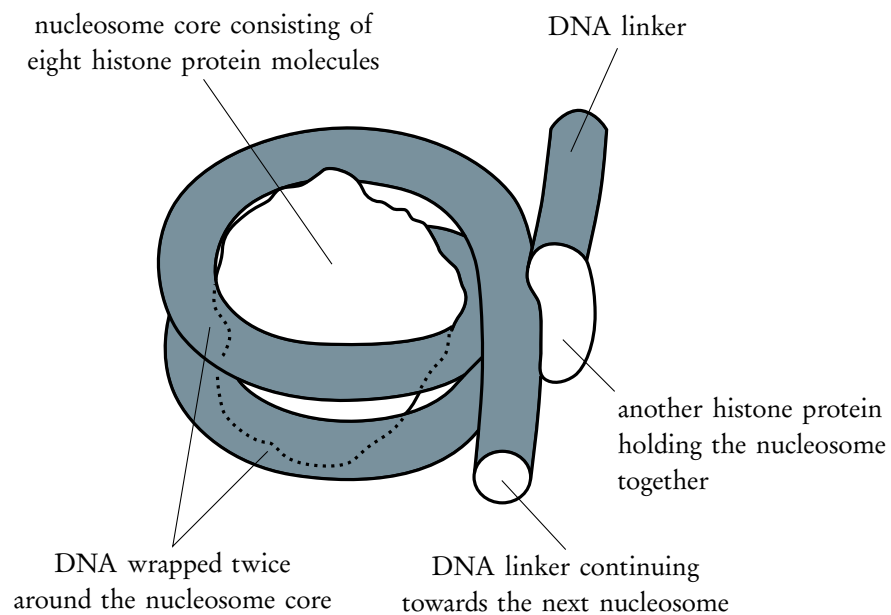


Figure 7.1: Structure of a nucleosome.

It is important to remember that histones are positively charged, while DNA is negatively charged. This makes the associations of these proteins with DNA very strong.

7.1.2 DNA replication

DNA consists of two antiparallel strands, both of which are in turn made of a sugar-phosphate backbone and one of 4 nitrogenous bases that attach the antiparallel strands through complementary hydrogen bonding:

Adenine (purine) forms two hydrogen bonds with Thymine (pyrimidine)

Guanine (purine) forms three hydrogen bonds with Cytosine (pyrimidine)

The binding specificity reduces the risk of mutations, as each nitrogenous base has a high binding affinity for its complementary base, and a lower one for the two remaining bases.

Next, an explanation of the replication process in prokaryotes at the two parent strands, also known as leading strand (where replication occurs continuously from the end of chromosome towards replication fork) and the lagging strand (where replication is not continuous and occurs from the replication fork towards the end of the chromosome):

1. Helicase unwinds and separates the DNA double-helix into the two parent strands, and DNA gyrase binds at the replication fork to stabilize the DNA during this process
2. Single strand binding proteins bind to the resulting parent strands to stabilize them and prevent them from reattaching and coiling once more

At lagging strand (3'–5' direction)

1. Primase attaches RNA primer is attached to begin replication at the site nearest to the fork, which contains the 3' end of the parent strand
2. As DNA continues to unwind and separate, more primers are attached at the replication fork
3. DNA polymerase III adds all the free nucleotides in a 5'–3' direction
4. DNA polymerase I replaces the RNA primer with DNA nucleotides
5. The separate segments that form between each primer Okazaki fragments
6. Enzyme DNA ligase joins Okazaki fragments together

At leading strand (5'–3' direction)

1. Primase attaches RNA primer is attached to begin replication at the 3' end of the leading strand
2. DNA polymerase III adds all the free nucleotides in a 5'–3' direction
3. DNA polymerase I replaces the RNA primer with DNA nucleotides

7.1.3 Non-coding DNA

It is important to realize that not all of the DNA present in the cell nucleus is later transcribed and translated to form proteins. Regions of DNA that code for proteins are called “exons” and regions that do not are called “introns.” Some special cases of non-coding DNA have very important functions in and outside the nucleus:

- Regulating gene expression: sites where proteins can bind to promote or repress transcription of specific genes
- Telomeres: sequences located on the tips of the chromosomes. After each replication event, a small segment at one end of the chromosome cannot be replicated, and that part of DNA is lost, thus leading to the shortening of telomeres (this way, important genes are not immediately lost)
- Genes for tRNAs and rRNAs: while these genes do not code for proteins, they do code for essential machinery needed during translation

7.2 Transcription and gene expression

7.2.1 Transcription

Transcription is the DNA directed synthesis of mRNA with RNA polymerase. As in DNA replication, the process of transcription, occurs in a 5′–3′ direction. RNA polymerase can only add free RNA nucleotides to the 3′ end of the growing mRNA molecule. Below is the description of the processes involved in transcription:

Initiation: RNA polymerase binds to the promoter region of the gene (this region contains a specific base sequence and signals the enzyme to initiate transcription)

Elongation: RNA polymerase simultaneously uncoils and transcribes the coding region in a 5′–3′ direction

Termination: RNA polymerase detaches from the mRNA strand after reaching the terminating sequence, the mRNA then “peels” away from the DNA template away from the DNA template and moves to be modified and later translated outside the nucleus

The DNA strand that is translated is called the “antisense” strand (the mRNA of interest will contain the sequence that is identical to the “sense” strand, which codes for genes of interest, with uracil instead of thymine).

7.2.2 Post-transcriptional mRNA modifications

In eukaryotic organisms, mRNA is modified before leaving the nucleus to be translated. The major modification consists in the removal of introns by RNA splicing, which “cuts out” the non-coding sequences and joins the remaining exons together. Many genes have several exons that can be spliced in various ways to create a large range of different proteins.

7.2.3 Gene expression: regulation of transcription

The process of transcription can be regulated by both proteins and environmental factors. Proteins that bind to the promoter region, known as transcription factors (TFs), are an example of this. TFs act by altering the ability of RNA polymerase to bind to the promoter and therefore affect the transcription of specific genes. There are normal TFs, activator TFs that enhance transcription, and repressor TFs that decrease the rate of transcription. Environmental factors can also affect gene expression in a way that is not inheritable. The study of epigenetics focuses on these factors and the way they alter transcription, mRNA modifications, translation, etc. For example, diet, age, and diseases can influence changes in gene expression that do not necessarily involve direct alteration of the DNA sequence. Some epigenetic modifications include:

- DNA methylation: can fix genes in an “off” position, preventing transcription
- Histone modification: acetylation and methylation can affect the level of chromatin condensation
- microRNAs: can bind to mRNA, leading to the degradation of the entire complex
→ gene cannot be translated

As mentioned earlier, the coiling and uncoiling of DNA by nucleosome interactions can regulate the process of translation. DNA regions of highly condensed chromatin will usually not be transcribed, as enzymes cannot easily access the DNA (regions that remain highly condensed at all times are known as heterochromatin, and are normally not transcribed). On the other hand, decondensed chromatin is easily accessible for transcription enzymes and transcription factors, and is readily transcribed.

7.3 Translation

7.3.1 Translation



Translation is the process in which mRNA transcripts are converted into polypeptide chains that can then be modified to create functional proteins.

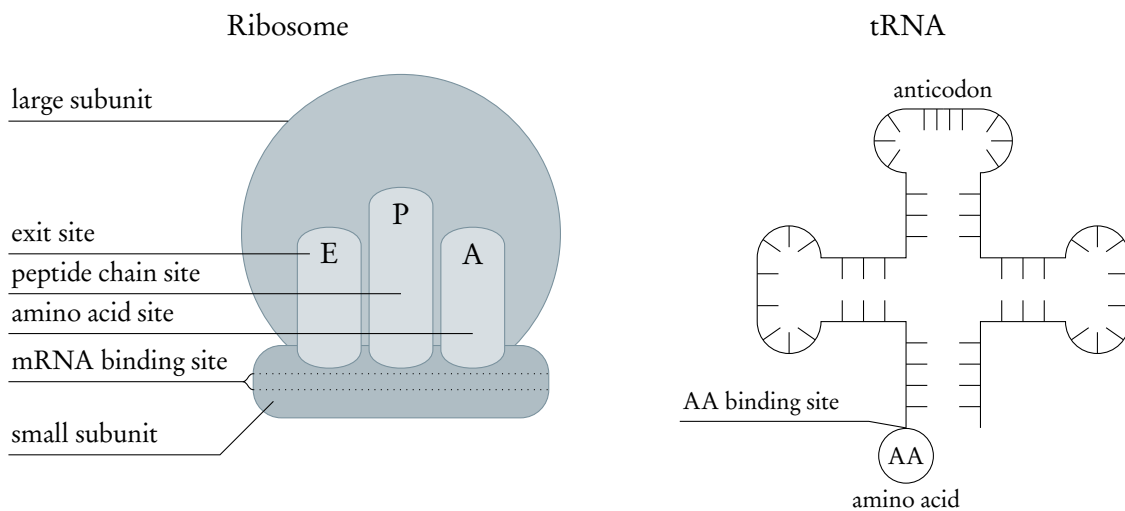


Figure 7.2: Ribosome structure and tRNA.

The process of translation is divided into three main stages and termination:

Initiation

The tRNA molecule with the start anticodon UAC binds to the small subunit of the ribosome (peptide chain site on diagram).

The small subunit then binds the 5' end of the mRNA strand and slides along it in a 5' to 3' direction until it finds the start codon on the mRNA (AUG).

The large and small subunit of the ribosome then bind together.

A tRNA molecule with the anticodon complementary to the next mRNA codon on the chain binds to the amino acid site of the ribosome.

The elongation process begins.

Elongation

The incoming tRNA pairs with its complementary mRNA codon at the amino acid site of the ribosome.

The amino acids in the tRNAs at the peptide and amino acid sites form a peptide bond, and the growing polypeptide chain remains attached to the tRNA at the amino acid site.

Peptide bond formation is catalyzed by the ribosome.

Translocation

The large subunit of the ribosome moves forward and the small unit slides after it, moving the entire ribosome 3 nucleotides down the mRNA strand.

The tRNA in the peptide site moves to the exit site (where the tRNA is released and can attach a new amino acid).

The tRNA carrying the growing polypeptide chain now moves to the peptide site.

The next codon to be translated is at the amino acid site, where its complementary tRNA codon can bind.

The process repeats itself.

Termination

When one of three stop codons (UAG, UAA, UGA) reaches the amino acid site of the ribosome, the large subunit advances over the small subunit, detaching from it and causing the polypeptide chain to be released from the last tRNA.

The polypeptide can then fold and be further modified to create a functional protein.

It is important to realize that translation of a single mRNA strand can occur in multiple ribosomes (many can attach), creating a polysome. Polysomes can be visualized using molecular visualization software.

It is important to realize that each tRNA molecule can only bind a specific amino acid, depending on its anticodon (triplet of bases located on the middle loop of the tRNA).

The amino acid binds at the 3' end of the single stranded molecule

20 different amino acids, some of which can bind to several tRNAs (same amino acid, different anticodons)

Skill: Identification of polysomes in electron micrographs of prokaryotes and eukaryotes → In the diagrams, you can clearly see the thin mRNA strand with multiple ribosomes attached to it, each translating the same mRNA (the darker circles show the growing polypeptide chains at each ribosome)

This binding is catalysed by tRNA-activating enzymes, which use the principle of substrate-enzyme specificity to attach the correct amino acid to the appropriate tRNA molecule

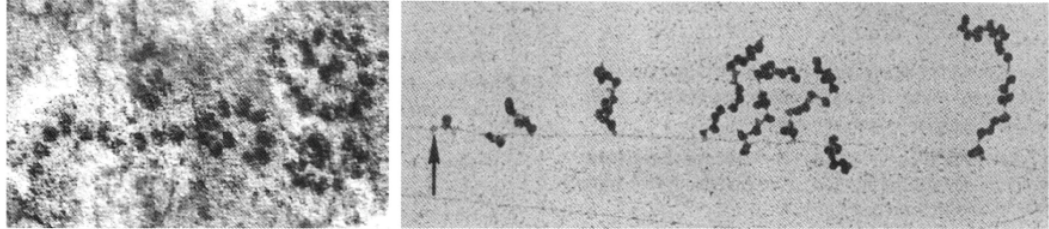


Figure 7.3: Polysome electron micrograph.

7.3.2 Free vs. bound ribosomes

In prokaryotic organisms, free ribosomes can begin the process of translation immediately after transcription, as there is no nuclear membrane. In eukaryotes however, the mRNA must first leave the nucleus through nuclear pores and can then be translated at either free cytoplasmic ribosomes, or bound ribosomes attached to the rough endoplasmic reticulum.

Free ribosomes release the protein into the cytoplasm, and the proteins formed by these organelles are mostly for use inside the cell (e.g., transcription factors, intracellular enzymes, etc.)

Bound ribosomes release the polypeptide chain into the rough endoplasmic reticulum, where the protein is modified, folded and packaged into vesicles that can then be secreted out of the cell or taken up by lysosomes.

To review the 4 levels of protein organization, refer to the section “Amino acids: the building blocks of proteins” in Chapter 2.

METABOLISM, CELL RESPIRATION AND PHOTOSYNTHESIS

8.1. Metabolism

162

- Metabolic pathways: chains and cycles
- Enzymes and activation energy of reactions
- Competitive vs. non-competitive inhibition
- End-product inhibition

8.2. Cell respiration

165

- Cell respiration: a redox reaction
- Glycolysis: from glucose to pyruvate
- Link reaction and Krebs cycle: aerobic cell respiration
- Electron transport chain and chemiosmosis: ATP
- Mitochondria: site for cell respiration

8.3. Photosynthesis

172

- Chloroplasts: the site for photosynthesis
- Light-dependent reactions
- Light-independent reactions

8.1 Metabolism



Metabolism is the sum of all chemical reaction occurring in a system (e.g. a cell or an organism). Enzymes are responsible for coordinating and controlling metabolic processes.

8.1.1 Metabolic pathways: chains and cycles

Several metabolic pathways consist of chains of reactions (e.g., glycolysis), where the initial substrate and end product do not necessarily interact or resemble one another. Other metabolic pathways consist of cycles of reactions (e.g., Krebs cycle), where a substrate of the cycle is constantly regenerated from other intermediates involved in the same cycle. Many metabolic pathways include both kinds of reactions.

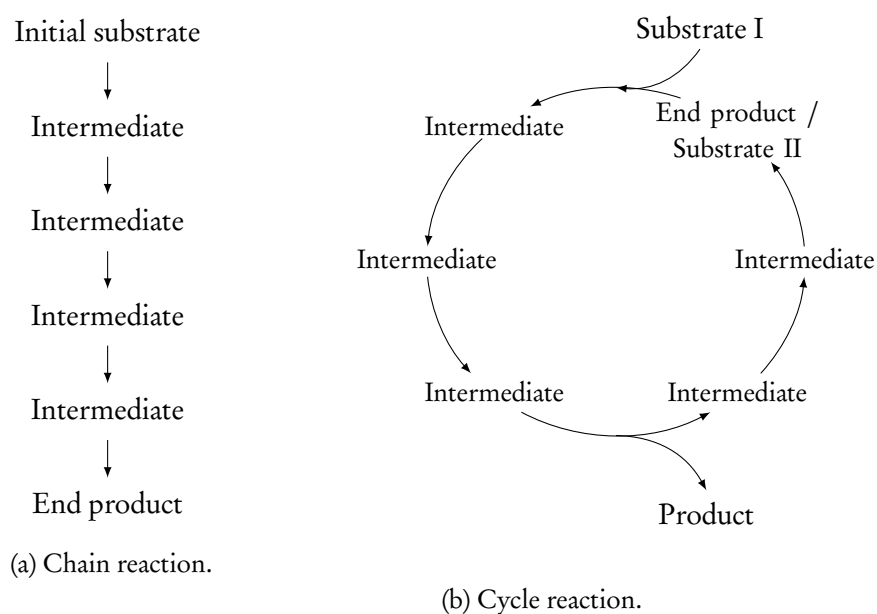


Figure 8.1: Diagram showing a chain and a cycle of reactions.

8.1.2 Enzymes and activation energy of reactions

The way enzymes catalyze, and thus speed up, reactions is as follows:

Before a molecule of the reactant can take part in the reaction, it has to gain energy (also known as activation energy), which helps break the molecular bonds within the reactant.

When new bonds are made after the reactant is broken down, energy is released (see Figure 8.2).

Enzymes reduce the activation energy of the reactions they catalyze by weakening the reactant's molecular bonds, making it easier for the reaction to occur.

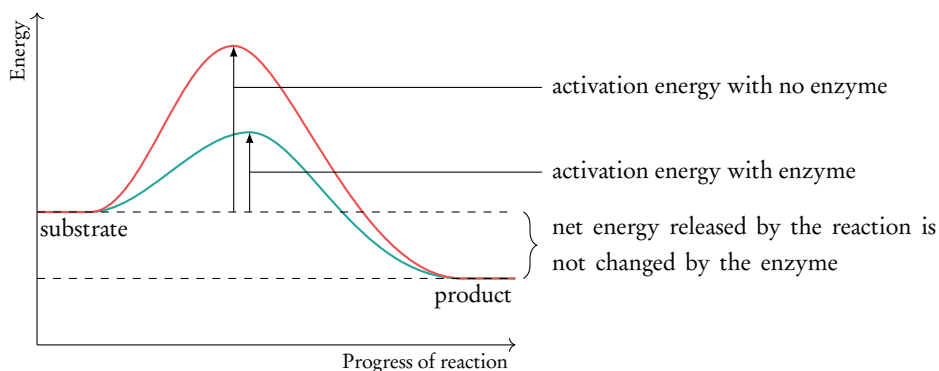


Figure 8.2: Graph showing the reduction in activation energy in the presence of enzymes, as well as the fact that the energy released by the formation of new bonds remains unchanged. The reaction shown is exothermic, meaning that it releases more energy than its activation energy (opposite is endothermic).

8.1.3 Competitive vs. non-competitive inhibition

Competitive inhibition

A competitive inhibitor competes directly for the active site of the enzyme by binding to it. It prevents the substrate from binding until the inhibitor dissociates. The inhibitor must have a very similar molecular structure as the substrate in order to fit the active site and be able to bind to it.

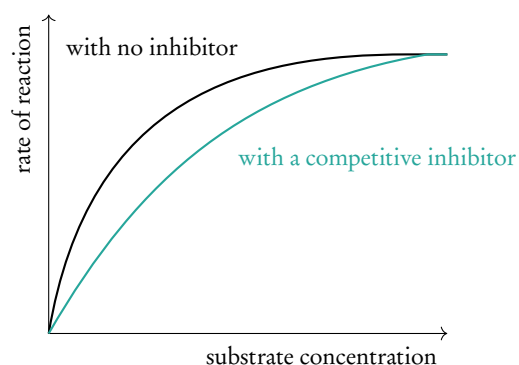
Competitive inhibition can be reversible or irreversible.

Increasing substrate concentration can usually reduce the inhibitory effect, as the inhibitor is more readily displaced by the many substrate molecules.

Example.

In the Krebs cycle, succinate dehydrogenase acts on substrate succinate, removing a hydrogen atom and converting it into another intermediate.

Malonate acts as a competitive inhibitor, effectively slowing down the Krebs cycle.



Non-competitive inhibition

A non-competitive inhibitor binds to the enzyme at a different site from the active site (also known as allosteric site).

Binding of the inhibitor at the allosteric site causes a conformational change in the enzyme (active site included), making it non-functional and slowing down the reaction.

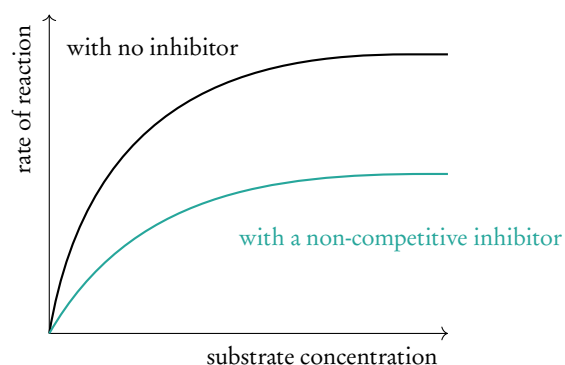
Non-competitive inhibition is usually, but not always, reversible but, unlike competitive inhibition, increasing substrate concentration does not reduce the inhibitory effect (even if there are more substrate molecules, they cannot bind to the non-functional active site).

Skill: Distinguish between different types of inhibition from graphs at specified substrate concentration.

Example.

Nerve gas (Sarin) binds to respiratory enzyme acetyl cholinesterase allosterically, changing the shape of the enzyme's active site.

Substrate can no longer bind, and this disruption can result in a blockade of the respiratory pathway → can be lethal, as this process is irreversible



8.1.4 End-product inhibition

End-product inhibition, also called feedback inhibition, occurs when the product of the last reaction in a metabolic pathway (chain or cycle) inhibits the enzyme that catalyzes the first reaction.

The inhibited enzyme is called the allosteric enzyme. Allosteric enzymes have both an active site and an allosteric site.

The end product binds to the allosteric site and changes the shape of the active site, inhibiting the enzyme, this process is reversible, so when the end product detaches, the reaction continues. This is an efficient way of controlling metabolic rate. If the concentration of the product is too high, end-product inhibition stops the pathway, producing less waste.

8.2 Cell respiration

8.2.1 Cell respiration: a redox reaction



Cell respiration is the controlled release of energy, in the form of ATP, from organic compounds in cells.

Cell respiration is a redox reaction, which involves the oxidation and reduction of electron carriers NAD^+ and FAD at different stages of this metabolic process. Electron carriers transfer chemical energy through redox reactions. Oxidation is the loss of electrons and hydrogen, whereas reduction is the gain of electrons and hydrogen.

Table 8.1

Oxidation	Reduction
Loss of electrons	Gain of electrons
Gain of oxygen	Loss of oxygen
Loss of hydrogen	Gain of hydrogen
Results in many C—O bonds	Results in many C—H bonds
Results in a compound with lower potential energy	Results in a compound with higher potential energy

8.2.2 Glycolysis: from glucose to pyruvate

Glycolysis is a chain reaction pathway that is catalyzed by enzymes in the cytoplasm of the cell and converts a glucose into two pyruvates. It uses no oxygen, so it happens in aerobic and anaerobic cell respiration (in both prokaryotes and eukaryotes). It is divided in four main phases:

Activation / Phosphorylation phase

Two molecules of ATP are used to begin glycolysis. One phosphate from each ATP phosphorylates the glucose molecule to form intermediate molecule fructose biphosphate

Lysis phase

This 6-carbon intermediate splits into two 3-carbon sugars, each containing one phosphate group, because it is very unstable.

Oxidation phase

Two hydrogen atoms are removed by NAD^+ from each 3-carbon molecule (oxidation). The energy released by this oxidation is used to link another free phosphate group to each 3-carbon molecule. The hydrogen atoms are then carried by electron carrier (forming $\text{NADH} + \text{H}^+$)

ATP formation phase

Pyruvate is formed by removing two phosphate groups from each 3-carbon molecules and phosphorylating ADP (yielding 4 ATP molecules). Because 2 ATP were initially used in the activation phase, glycolysis yields the following net products:

- $2 \times \text{ATP}$
- $2 \times \text{pyruvate}$
- $2 \times \text{NADH} + \text{H}^+$

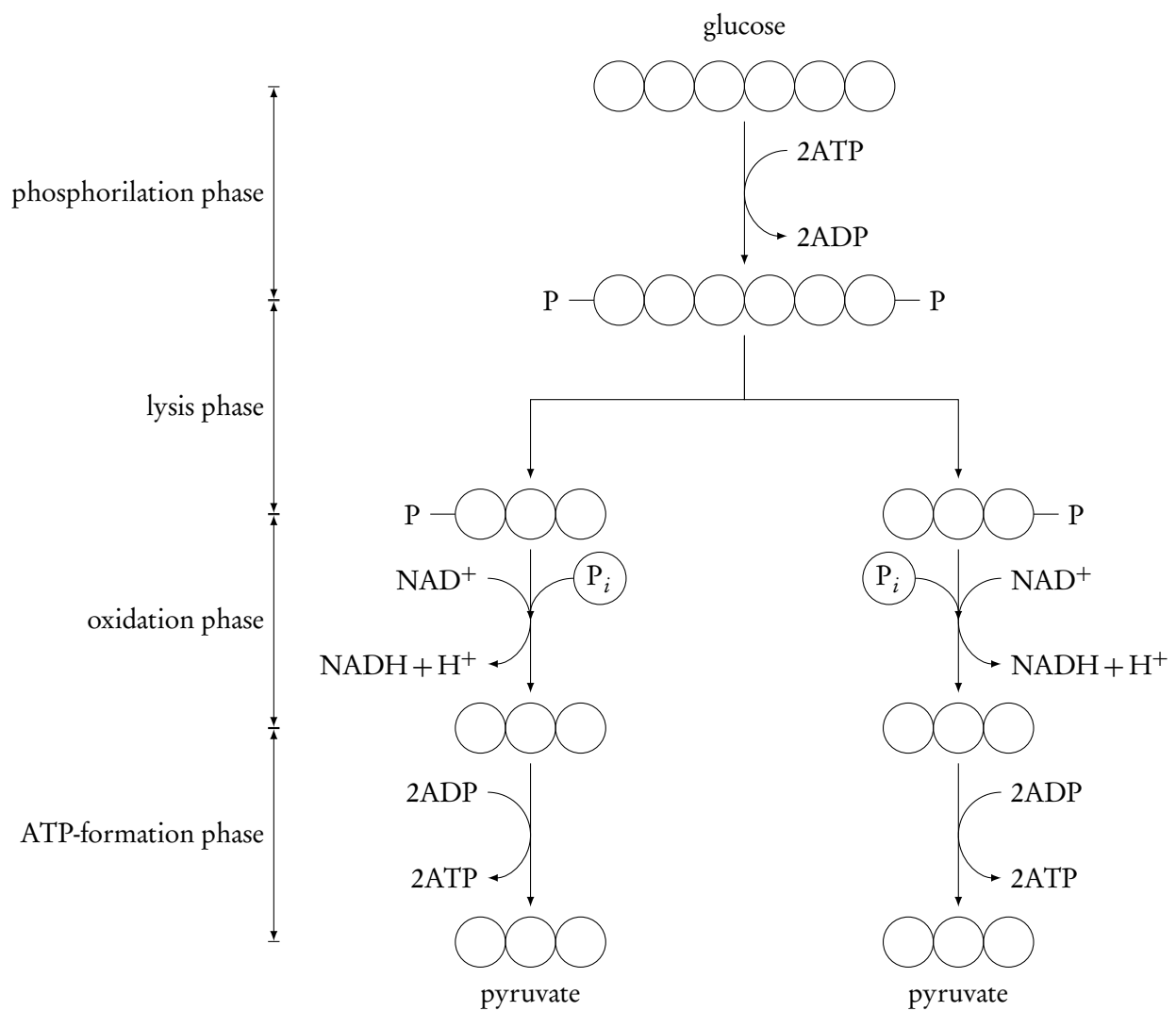


Figure 8.3: Glycolysis.

8.2.3 Link reaction and Krebs cycle: aerobic cell respiration

If oxygen is readily available to the cell, the pyruvate molecules undergo the link reaction and the Krebs cycle to create a much higher yield of ATP. Next, a description of both processes.

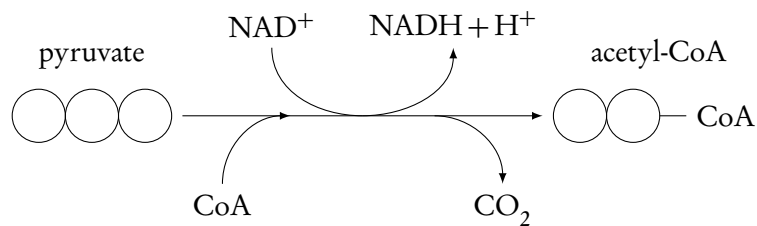
Link reaction

Pyruvate enters the mitochondrion's matrix through active transport. In the matrix, each pyruvate undergoes the following steps:

- Oxidation by NAD^+
- Decarboxylation
- Binding to coenzyme A (CoA) to form acetyl-CoA

Link reaction yields:

- $2 \times \text{CO}_2$
- $2 \times \text{NADH} + \text{H}^+$
- $2 \times \text{acetyl-CoA}$



Krebs cycle

Occurs in the matrix of the mitochondrion.

Acetyl-CoA combines with a 4-carbon compound, forming a 6-carbon intermediate (all the products mentioned below double, since there are two acetyl-CoA molecules that enter the cycle).

The 6-carbon intermediate undergoes oxidative decarboxylation, resulting in a 5-carbon molecule, CO_2 and $\text{NADH} + \text{H}^+$.

Another oxidative decarboxylation occurs, resulting in a 4-carbon molecule, CO_2 and $\text{NADH} + \text{H}^+$.

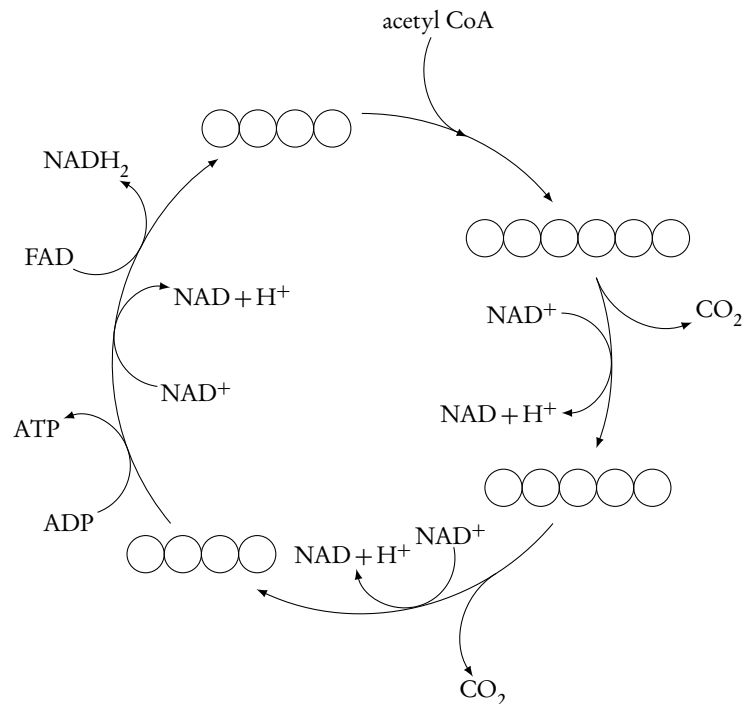
The 4-carbon intermediate undergoes substrate-level phosphorylation, releasing a phosphate group to phosphorylate an ADP molecule. The resulting intermediate is then oxidized, this time by $\text{FAD} \rightarrow \text{FADH}_2$.

The resulting 4-carbon molecule is the same as the starting intermediate that combines with acetyl-CoA at the beginning of the cycle.

Overall, the two acetyl-CoA molecules lead to the following products:

- $2 \times \text{ATP}$
- $6 \times \text{NADH} + \text{H}^+$
- $2 \times \text{FADH}_2$
- $4 \times \text{CO}_2$

CO_2 is always released as a waste product (in total 6 molecules) in cellular respiration.



8.2.4 Electron transport chain and chemiosmosis: ATP



Chemiosmosis is the movement of ions across a semipermeable membrane, down their electrochemical gradient.

Through glycolysis and the krebs cycle, several electron carriers high in energy are produced. $\text{NADH} + \text{H}^+$ and FADH_2 are responsible for carrying carry hydrogen and electrons (high in energy) from the mitochondrial matrix to the mitochondrial inner membrane. NAD^+ allows the production of 3 ATP molecules and FAD the production of 2 ATP molecules upon entering the electron transport chain. Next, a description of the events that take place at the mitochondrial inner membrane, which lead to the largest yield of ATP: the electron transport chain and chemiosmosis.

Electron transport chain and chemiosmosis

Electron carriers are located on the inner mitochondrial membrane, each can receive electrons at different energy levels.

$\text{NADH} + \text{H}^+$ and FADH_2 are oxidized, releasing electrons to the carriers and H^+ into the matrix.

Electrons “jump” from one carrier to the next, with each jump releasing some energy as the electron goes to a more energetically favorable position.

The energy released allows for the hydrogen ions at the matrix to be pumped into the small intermembrane space, against their concentration gradient.

At the end of the carrier chain, de-energized electrons return to the matrix and join with oxygen and hydrogen atoms, forming water as a waste product.

At the intermembrane space, hydrogen ions accumulate at a high concentration. This creates both a chemical gradient of hydrogens (high at the intermembrane space and low at the matrix) and an electrical gradient (more positive in the intermembrane space and more negative at the matrix), thus forming the electrochemical gradient)

ATP synthase is an enzyme found embedded in the inner membrane of the mitochondria. With the electrochemical gradient of H^+ pointing toward the matrix, H^+ ions naturally go into the ATP synthase. When this happens, the enzyme can use the energy released by the H^+ going into a more energetically favorable state (going to the matrix) to join ADP with P_i to form ATP. With the charged electron carriers yielded from one glucose molecule, ATP synthase has enough energy to yield a large number of ATP molecules, between 36 and 38.

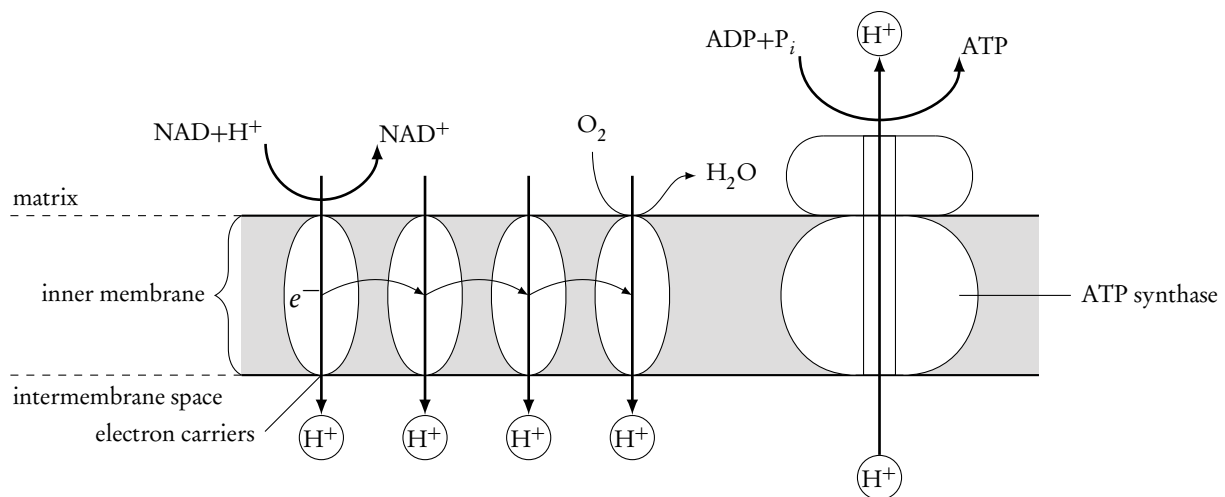


Figure 8.4: Electron transport chain.

Therefore, in the presence of oxygen cells are able to produce much more ATP (36–38) than in anaerobic conditions (4).

8.2.5 Mitochondria: site for cell respiration

Mitochondria are commonly referred to as the “powerhouse” of the cell. This organelle is where the link reaction, Krebs cycle, electron transport chain and chemiosmosis occur, making it essential for providing the cell with the energy needed to meet its functional demands. The structure of mitochondria is very advantageous for the various functions it must accommodate. Figure 8.5 is an annotated diagram:

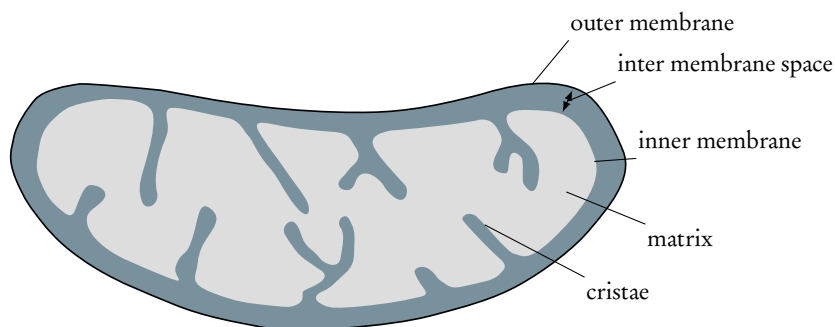


Figure 8.5: Mitochondria.

Skill: Annotation of a diagram of a mitochondrion to indicate the adaptations to its function.

Outer membrane: separates the contents of the mitochondrion from the rest of the cell

Inner membrane: contains the electron transport chain and ATP synthase, which carry out oxidative phosphorylation.

Intermembrane space: this small space between mitochondrial membranes allows for the rapid accumulation of protons (H^+) necessary for chemiosmosis to occur

Cristae: tubular regions surrounded by membranes that increase surface area for oxidative phosphorylation and the electron transport chain

Matrix: internal cytosol-like fluid that contains the enzymes necessary for the Krebs cycle and link reaction

8.3 Photosynthesis



Photosynthesis is the process used by plants to produce their own organic substances (most commonly glucose) using light energy from the Sun and simple organic substances.

Photosynthesis involves multiple reactions that can be categorized into light-dependent and light-independent. Light-dependent reactions produce intermediate compounds that are later used in the light-independent reactions to produce organic “fuels” for the plant.

8.3.1 Chloroplasts: the site for photosynthesis

Chloroplasts, organelles that are unique to plant cells and eukaryotic algae, are the site where photosynthesis reactions take place. Their structure has evolved for their function as can be seen in the annotated diagram in Figure 8.6.

Skill: Annotation of a diagram of a chloroplast to indicate the adaptations to its function.

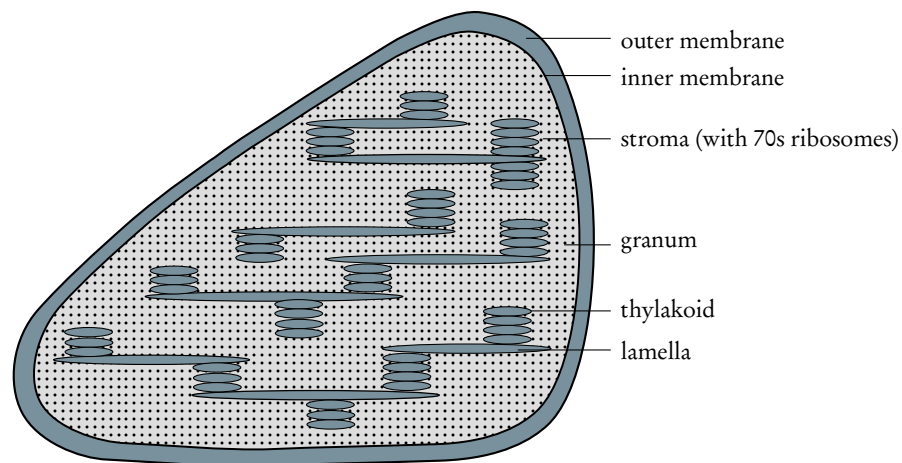


Figure 8.6: Chloroplasts.

Thylakoids: the small, disc-shaped structures increase the surface area for light absorption by photosystems I and II

Thylakoid space: the small space inside thylakoids allows for more rapid accumulation of protons to create an electrochemical gradient for chemiosmosis

Stroma: cytosol-like substance where all the enzymes needed for the Calvin cycle are found

Double membrane: isolates the working parts of enzymes within the chloroplasts from the surrounding cytosol

Lamella: connect granules and further increase surface area for light absorption

8.3.2 Light-dependent reactions

Light-dependent reactions take place at the thylakoid membrane, where photosynthetic pigments are concentrated in photosystems I and II (PSI and PSII). The light-dependent reaction can be divided into 5 main steps:

1. Light from the Sun is absorbed by chlorophyll in PSII. This causes photolysis, the splitting of water by light energy into oxygen (released as waste), hydrogen (protons), and excited electrons that are transferred to PSII.
2. Electrons in PSII are excited by light from the sun and “jump” to a higher energy level, where they are received by an electron acceptor → this is called photoactivation of PSII.
3. These electrons get transferred to an electron transport chain within the thylakoid membrane. As they go down the electron transport chain they release energy that allows the pumping of protons from the stroma into the thylakoid space. Just like in cellular respiration, the pumping of protons into the thylakoids creates an electrochemical gradient rich in energy.
4. Proton accumulation allows chemiosmosis to take place with the production of ATP through ATP synthase. Because ADP phosphorylation depends on the use of light (photons), this process is referred to as non-cyclic photophosphorylation.
5. De-energized electrons are taken up by photosystem I (PSI), and are once again excited due to the absorption of light and taken up by an electron acceptor. From here, two things may occur:
 - Electrons enter the electron transport chain once more, to promote further yield of ATP in a process referred to as cyclic photophosphorylation.
 - If ATP levels are high enough, electrons are taken up by electron carrier NADP to form NADPH, and carried away to the stroma to be used in the light-independent reactions.

Overall, the light-dependent reactions produce:

- ATP
- NADPH
- O₂ (released as a by-product)

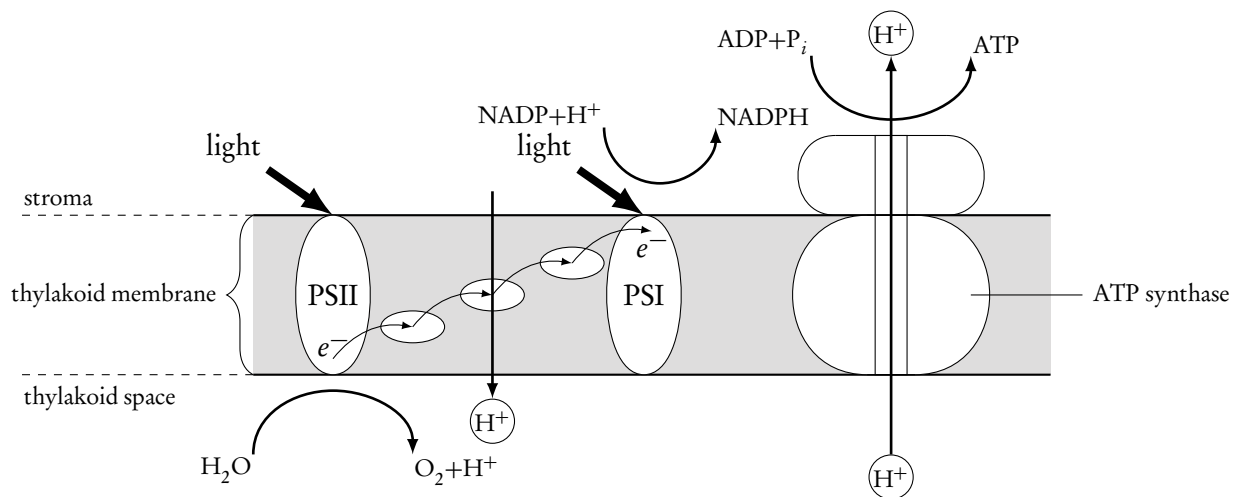


Figure 8.7: Light-dependent reactions

8.3.3 Light-independent reactions

The light-independent reaction (or Calvin cycle) is characterized by the carboxylation of ribulose biphosphate to produce glucose and other organic compounds. It occurs at the stroma of the chloroplast, where the products of the light-dependent reaction (ATP and NADPH) as well as carbon dioxide, that has diffused into the chloroplast, can be found.

The cycle begins and ends with 3 5-carbon molecules called ribulose bisphosphate (RuBP). There are 3 main steps that outline the Calvin cycle:

1. **Carbon fixation:** the 3 RuBP molecules undergo carbon fixation with the aid of enzyme rubisco, which catalyzes the molecules' carboxylation using three carbon dioxide molecules. This yields 3 6-carbon intermediate molecules that are very unstable, and thus immediately split into 6 3 carbon molecules of glycerate-3-phosphate (G3P).
2. **Reduction:** all G3P molecules are reduced by NADPH and phosphorylated by ATP, yielding 6 3-carbon triose phosphate (TP) molecules.
3. **Regeneration of RuBP:** One TP molecule leaves the cycle and is used to make half of a glucose molecule. Two cycles of the Calvin cycle are thus needed to make one glucose molecule. The remaining five TP molecules are phospholyrated by ATP and rearranged back into the intial 3 RuBP molecules the cycle started with.

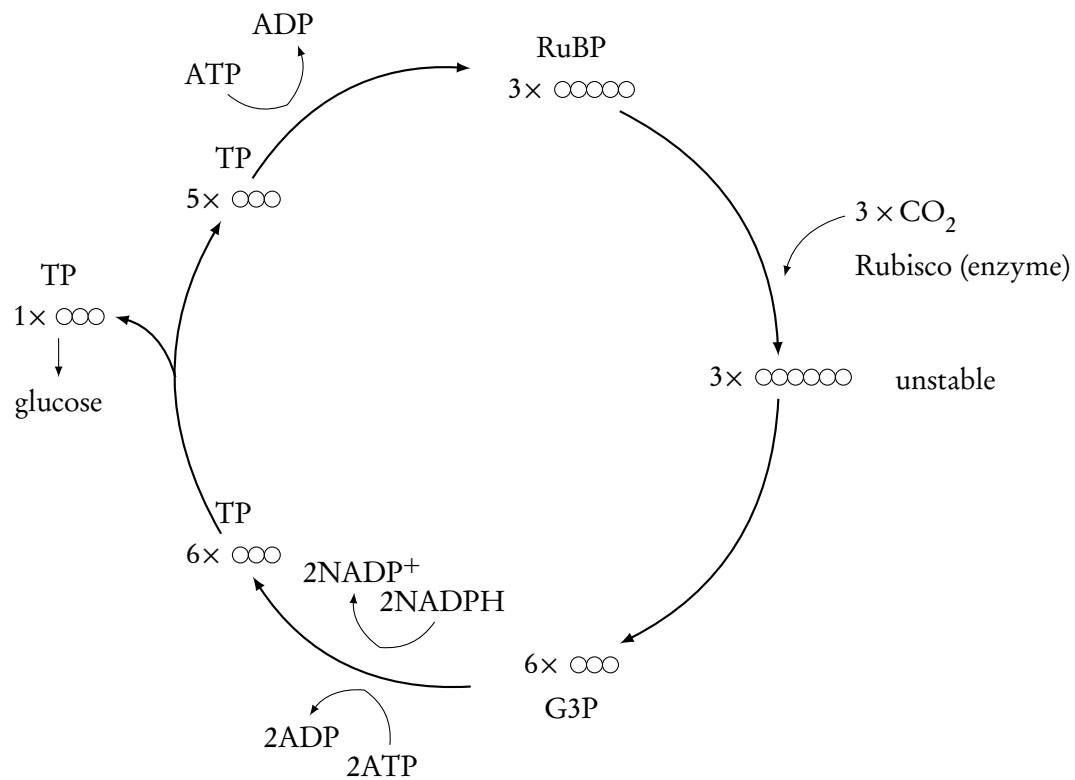


Figure 8.8: Calvin cycle.

9.1. Transport in xylem **178**

- Water transport
- Adaptations of plants to different environments
- Factors affecting transpiration rate

9.2. Transport in phloem **184**

- Structure
- Function
- Hydrostatic pressure
- Co-transporter
- Measuring the rate of phloem transport

9.3. Reproduction of flowering plants **186**

- Flowering and day length
- Mutualism and pollination

9.4. Growth in plants **188**

- Growth of the shoot
- Auxin and plant growth
- Germination
- Micropropagation

9.1 Transport in xylem



Transpiration is the continuous movement of water from the roots of a plant to the leaves where water is lost as water vapour. Due to the adhesive and cohesive properties of water, the loss of water at the leaves creates a constant pull/movement of water through the plant.

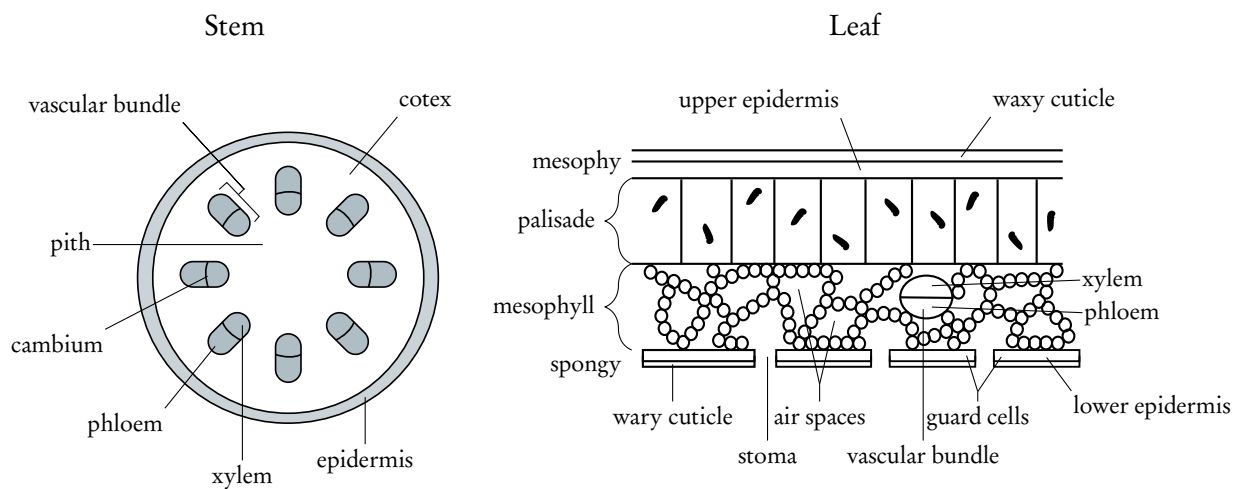


Figure 9.1: Leaf structure.

Palisade mesophyll is the main photosynthetic tissue, and makes up the top part of the leaf, rich in chloroplasts.

Spongy mesophyll is the loosely packed mesophyll that provides the surface for gas exchange. It faces the bottom side of the leaf.

Stoma are small gaps in the lower epidermis that allow gas exchange. These can open and close with the help of guard cells to regulate/minimize water loss.

Waxy cuticle prevents excessive water loss from both the top and bottom side of the leaf.

Epidermis (both upper and lower), produces the waxy cuticle and prevents water loss.

In general, the inside of the leaf needs to be as moist as possible to allow good gas diffusion from and to the palisade mesophyll. Even though the leaf is designed to reduce water loss as much as possible, the stoma allow for escape of some water vapour from the plant leaves.

Xylem and phloem are two vessel structures in a plant that ensure that all parts of the plant are supplied with water and nutrients.



Xylem is the main water transport vessel and can be recognized in a picture by its large hollow spaces.

Phloem is the main vessel for transport of organic molecules, such as sugars and amino acids through a sap.

Cambium makes up the interface of xylem and phloem and can become either one of them.

9.1.1 Water transport

Root

The main part of water absorption occurs in the root of the plant. The soil contains a lot of water, but also mineral ions.

With the large surface area of the root, the plant is able to absorb a lot of water from the soil by osmosis, as the root cells are more saturated in sugars and ions compared to the soil.

The mineral ions can only be absorbed through active transport since the concentration of ions is greater in the root than in the soil (think about the volume of a root compared to the volume of the total soil in a forest - which is more easily saturated?).

There are mitochondria and protein pumps in the root cells that allow the plant to take up sodium, potassium and other minerals through active transport.

Stem

Xylem is the water transporting vessel in the plant.

It is a tubular structure, made of dead plant cells that have fused together.

The primary xylem originates in the root and is strengthened by a thick cell wall and lignin.

The rest of the xylem is also often impregnated with lignin which helps the xylem cells resist the inward collapse that would occur in very low pressures.

Water is pulled from the roots to the top of the plant through transpiration. The mass flow of water is achieved due to three main properties:

1. Water pressure is high at the roots (water in) and low at the leaves (water out) creating a pressure gradient.
2. Adhesive property of water allows water molecules to adhere to the xylem wall and move upwards.
3. Cohesive property of water allows water molecules to stick to one another, creating a continuous stream.

Recall that the cohesive and adhesive properties of water allow it to make a continuous stream and adhere to the walls of xylem (respectively).

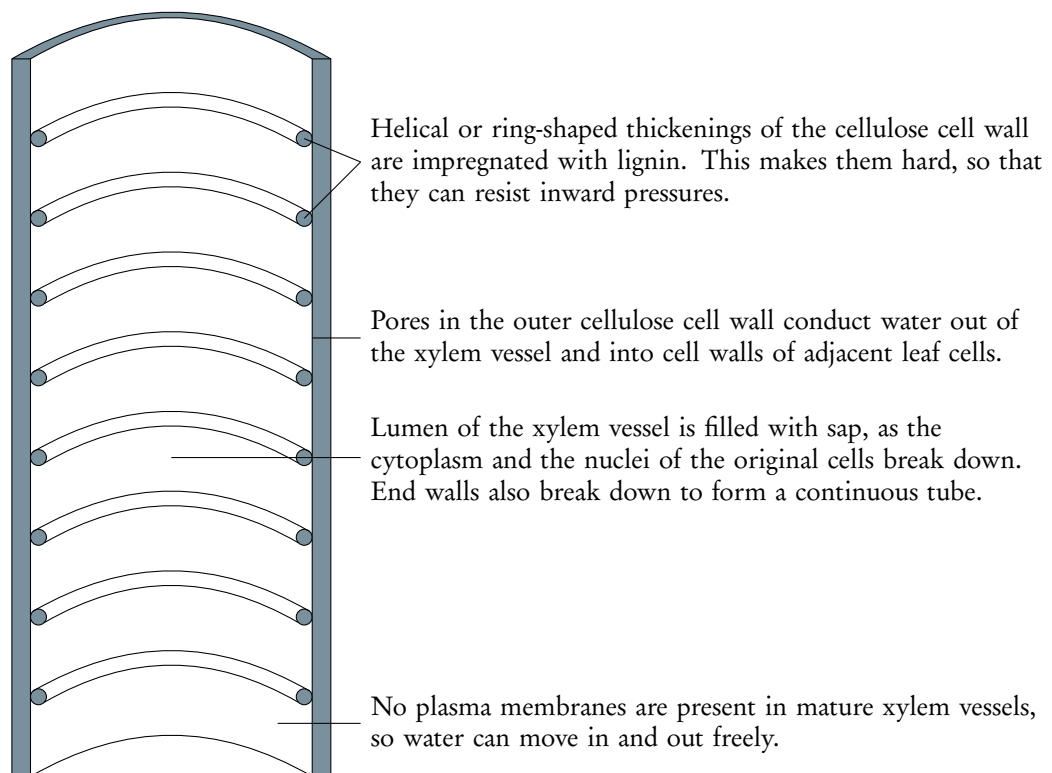


Figure 9.2: Primary xylem vessel.

Xylem cells are dead which allows movement of water. Lumen is filled with water. The ring shaped structures are thickenings of cell wall surrounded by lignin. Pores in the outer cell wall allow for water to move in and out of the xylem to the surrounding tissues.

There are three main assumptions about water transport that can be tested by simple experiments:

1. Water has adhesive properties
2. The cell wall contains small capillaries through which the water moves
3. Evaporation of water leads to tension in the xylem

Experiments that test these three assumptions:

1. If a glass capillary (hollow) is placed in a bowl of water, the water will slowly be pulled up the capillary. This is not the case if you replace water by mercury which doesn't have adhesive properties.
2. If you dip a strip of paper in water (and paper is made of cellulose), the water will slowly move up the strip and you will see more and more of the paper getting wet
3. If you take a pot with many small holes, seal it and put a small tube with water excess through it, you will see the water gradually rising through the tube, because the water vapour in the pot will slowly evaporate through the holes and create tension inside the pot

9.1.2 Adaptations of plants to different environments

Saline soil

In areas with dry climates, water evaporates from the soil easily and the soil is left saturated with mineral ions.

Due to high solute concentration in such soil, the water would not be able to move through osmosis into most plants.

Halophytes are plants adapted to saline soils as they keep the solute concentrations inside their roots, higher than the concentration of the soil.

They do this by keeping high concentration of potassium and sugars in their cytoplasm (not sodium since high sodium impairs metabolic functions).

In the vacuoles, they can keep high levels of Na^+ since there are no metabolic processes going on there. Moreover, there is active excretion of sodium out of the plant.

In this way, halophytes maintain high ion concentrations so that the little water from the soil still enters the plant by osmosis.

Desert

Xerophytes are plants adapted to life in dry habitats. Cacti are examples of xerophytes and they thrive in deserts.

There are several adaptations that will help them achieve this:

- They have a thick waxy cuticle that prevents excessive water loss.
- Their stems are very upright, so that the largest parts of the stems are only exposed to sunlight in the morning and evening (when the sun is not straight above them) when the heat is less extreme.
- Their stomata have the so called CAM physiology, which allows them to close during daylight and open at night, which is the opposite of what happens in normal plants (recall that stomata are needed for gas exchange during photosynthesis that occurs during daylight).
- The areas of their leaves is reduced, so cacti have spines instead of true leaves to reduce water evaporation.



Potometer is a device designed to measure the rate of transpiration of plants

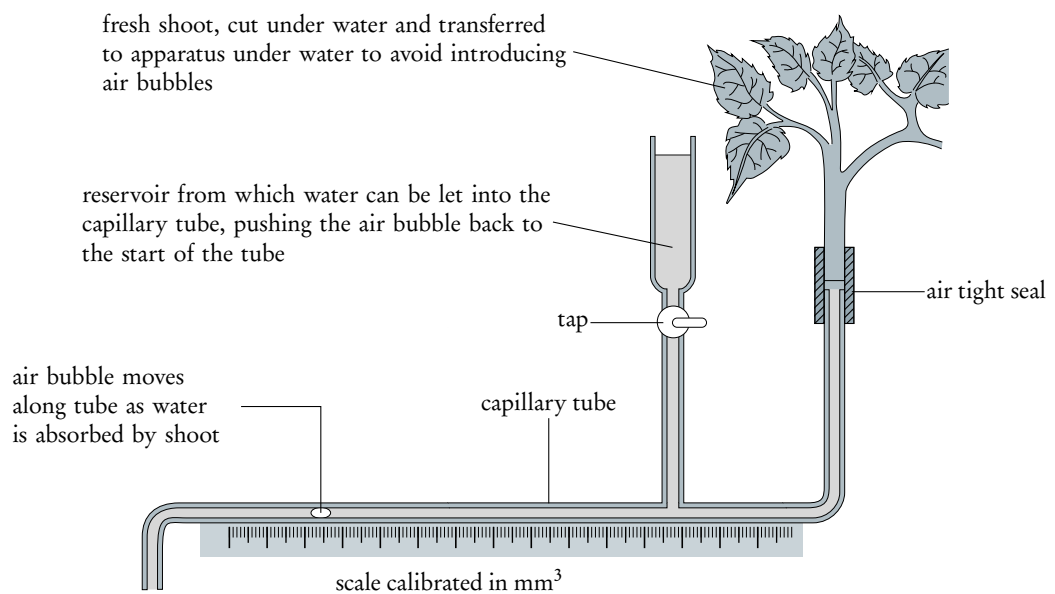


Figure 9.3: Potometer.

The plant is tightly attached to a long capillary tube that has a water source.

There is an air bubble inside the capillary that indicates how much water the plant has taken up.

As the water transpires off of the plant's leaves, the water is taken up into the capillary and the bubble moves.

In order to calculate a precise amount of water that was taken up, the measurements should be repeated several times.

9.1.3 Factors affecting transpiration rate

Temperature

At higher temperatures, the water molecules evaporate more easily from the surface of the leaves.

At higher temperatures, air molecules around the leaf move faster, so the relative humidity is also lower, creating a higher gradient for water to move out of the plant.

At too high temperatures, the leaves' stomata may close, and the transpiration decreases slightly.

How to measure this factor using a potometer?

Using a heat lamp directed at the plant, and a thermometer to measure leaf temperature, you can vary the temperature of the plant

Humidity

Humidity refers to the amount of water molecules in the air.

The higher the humidity, the smaller the concentration gradient between the moist mesophyll and humid environment, so the transpiration decreases.

How to measure this factor using a potometer?

- A plastic bag around the plant will make sure no air escapes
- By spraying moisturising spray in the bag, or by adding silica bags to the plastic bag, you can increase/decrease the humidity around the plant
- The relative humidity can be measure using a hygrometer

Wind

If the air around the leaf is still, the water vapour in the air will not move far away from the leaf and the humidity will increase.

The higher the wind speed, the lower the humidity around the leaf, and the higher the transpiration.

At too high wind speeds, the stomata may close.

How to measure this factor using a potometer?

- An electric fan can produce movement of air around the plant, and you can vary its velocity by moving the fan closer or further away from the plant
- An anemometer is used to measure the exact speed of air movement

9.2 Transport in phloem



Phloem is the vessel transporting organic compounds within the plant, a term often substituted by the term phloem sieve tube.

Source (in the phloem) constitutes photosynthetic tissue (e.g. leaves).

Sink (in the phloem) constitutes organs of storage of organic material (e.g. fruits, roots).

9.2.1 Structure

The phloem is made of live cells that break down their nucleus and most of the organelles.

The walls in between the cells contain many holes called the sieve plate, which both help maintain the structure of the tube, and allow for sap to pass through either direction.

The membrane of cells contains many protein pumps that load and unload sucrose.

The membrane also contains plasmodesmata, tight connections between the phloem cells and the companion cells in the vicinity to mediate exchange of materials..

9.2.2 Function

The phloem brings organic molecules to all parts of the plant.

The photosynthetic parts of the plant produce organic “food” molecules and are therefore called sources.

In the sources, sucrose is being loaded into the phloem through active transport. This is because the phloem tube is highly concentrated in organic molecules.

The storage parts of the plants, such as roots (carrots), tubers (potatoes) or fruits, are called sinks, as this is where the sucrose gets unloaded from the phloem. Companion cells exist between both the source and the sieve tube and between the source and the sieve tube. This is an intermediate location for the organic products.

9.2.3 Hydrostatic pressure

Hydrostatic pressure is the pressure exerted by a fluid and is maintained by different solute concentrations in the liquid and surrounding spaces.

In a plant, sources are rich in sugars causing osmosis, or the movement of water from the cells surrounding the source, and therefore the xylem, toward the sieve tube. This leads to an increase in hydrostatic pressure.

At the sources the solute concentrations are low at the sieve tube, as organic molecules are being stored away. Water can therefore leave the phloem and enter the surrounding tissues and eventually the xylem.

This leads to a lower hydrostatic pressure and therefore a pressure gradient between sieve tubes at sources and sinks. This allows for a constant movement of the sap at the sieve tube from the areas by the sources toward the area of the sink. In this way organic molecules are constantly and efficiently being moved from the site of production to the site of storage.

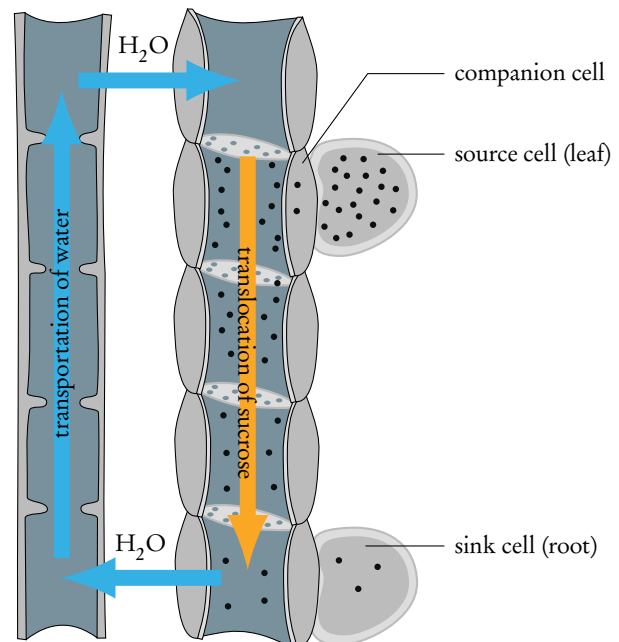
9.2.4 Co-transporter

Sucrose is loaded from the source into the phloem via active transport.

This active transport occurs with the help of co-transporter proteins that use the gradient of ions to move molecules like sucrose from the tissue to phloem.

One such co-transporter pumps sucrose into the phloem by making use of a hydrogen ion gradient that is created by a hydrogen protein pump. This is therefore a type of secondary active transport, where the co-transporter uses the energy stored in the hydrogen gradient to move sucrose against its concentration gradient. (recall that ATP synthase also uses the energy of the proton flow to make ATP).

Companion cells can help with this process as they take up the sucrose from the source, and then pass it onto the sieve tube as they have a highly concentrated environment.



9.2.5 Measuring the rate of phloem transport

Aphids are small animals that feed on the sap of plants.

They do this by piercing the plant with their stylets until they reach the phloem sieve tube.

If the scientist supply the plant with carbon dioxide containing radioactive carbon, the plant makes radioactive sucrose that can be detected in the phloem.

Once the aphid pierces the phloem, the stylet is cut off, and the sucrose-rich sap that comes out can be collected.

The time taken for the radioactive sucrose to move to different parts of the phloem can be used as a measure of phloem transport rate.

9.3 Reproduction of flowering plants

Flowers are the reproductive organs of many plants. The stamen constitutes the “male” organs and the carpel constitutes the “female” organs.

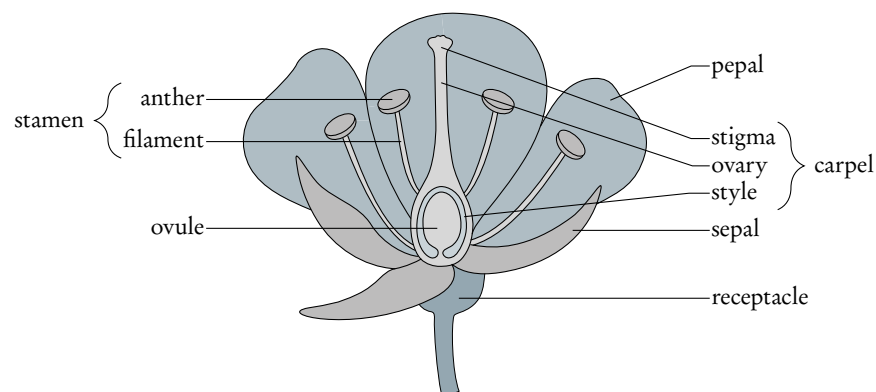


Figure 9.4: Flower.

The seed contains the embryo of a plant and supplies the food and water needed for its development. The seed is made of modified leaves called cotyledons.

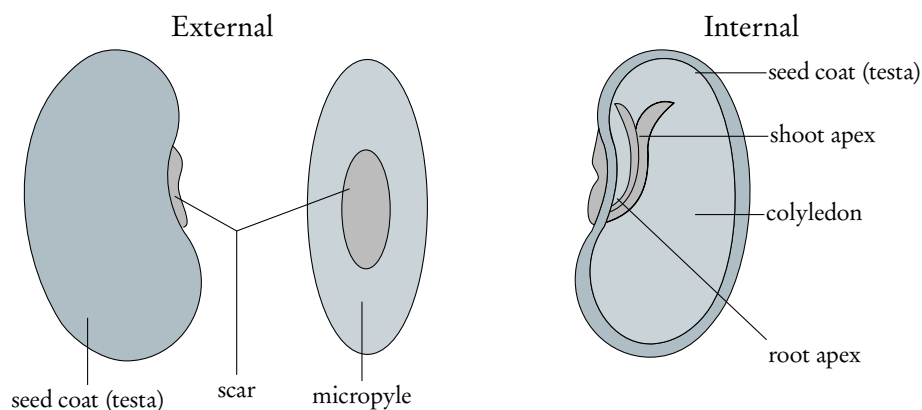


Figure 9.5: Seed.



Fertilization is the process of zygote formation, which occurs through the fusion of male gametes (in pollen) and female gametes (in ovule).

Pollination is the transfer of pollen (containing male gametes) from anthers to stigma (which contains the female gamete). Pollination occurs as the pollen is transferred to the stigma, where the gametes germinate and grow in a pollen tube to the ovary.

Seed dispersal is the process where the seeds containing the plant embryo are dispersed in nature where they germinate and grow. Seed dispersal occurs as the fertilised ovules develop into fruits that contain seeds.

Each of these three processes needs to occur for the plant to reproduce successfully!

9.3.1 Flowering and day length

Different plants flower at different times of the year.

Short day plants flower in winter, while long day plants flower in summer.

It is the length of the dark period that determines when the plant will flower. This is regulated by a molecule called phytochrome that exists in two forms:

- inactive red form (Pr): absorbs red light causing it to become Pfr.
- active far red form (Pfr): absorbs far red light causing it to become Pr.

Sunlight contains much more red light than far red light, therefore:

- during the day Pr gets converted into Pfr, reaching the highest levels of Pfr.

- at night, or in the absence of sunlight, Pfr reverts back to Pr.

Short and long day flowering plants depend on the levels of Pfr to determine when they will flower:

Short day plants (winter flowering): Pfr inhibits the flowering of the plant, therefore a longer night will cause the lowest amounts of Pfr, causing flowering.

Long day plants (summer flowering): Pfr triggers flowering, therefore shorter nights will cause low amounts of Pfr changing to Pr and an accumulation of Pfr, causing flowering.

Plants can be stimulated to flower “out of their season” by varying the amount of time when they are exposed to light vs. dark.

9.3.2 Mutualism and pollination

Most of the flowering plants depend on insects and other animals for reproduction.

This is called a mutualistic relationship, since the animal that pollinates the flower feeds on the nectar from the plant, and in return, it transfers pollen from one plant to another (or within one plant).

Often, one specific species of insect pollinates one specific plant.

An example of this is a species of bee that feeds on vanilla orchid nectar and therefore pollinates only vanilla orchids by crossing from one plant to another in search for food.

9.4 Growth in plants



Meristems are regions of plants containing undifferentiated cells that continuously divide and grow.

Determinate growth can be observed in animal species, where the embryo does not grow indefinitely, but has a determined number of legs, arms, organs, etc, to grow.

Indeterminate growth is seen in plants, where the apical meristem can continually provide new cells for further growth of the plant (e.g. meristems).

9.4.1 Growth of the shoot

Apical meristem are found in shoots and roots of the plant, and are responsible for lengthening of the plant (primary growth) and leaf development.

Shoot of the plant is defined as the stem with leaves. As mentioned earlier, the shoot contains the cells of apical meristem which continuously divide

During these divisions, some of the cells get displaced to the sides of the meristem and therefore stop dividing.

Instead, they grow and differentiate in order to produce stem and leaf tissues.

Leaves are produced when the meristem cells get displaced to the side of the apical meristem and form bumps.

These bumps, called leaf primordia, are differentiated cells that continue to divide and grow until they form full grown leaves.

Lateral meristem are found at the cambium of the plant, and are responsible for lateral growth (secondary growth) or thickening of the plant and production of bark in woody plants.



Auxin is a plant hormone that controls growth in the shoot tip, by stimulating or inhibiting cell division, and setting the direction of growth (tropism).

Phototropism is directional growth guided by the brightest source of light.

9.4.2 Auxin and plant growth

There are photosensitive pigments in plant shoot cells, called phototropins that respond to different light intensity.

Once activated, the photopigments also activate proton pumps that pump H^+ ions into the cell wall.

In the cytoplasm, auxin is a negatively charged molecule.

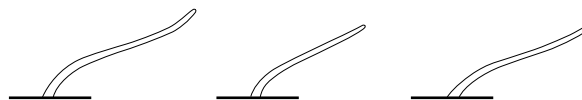
The positive charge of the wall attracts auxin molecules to move towards the wall, where they bind a hydrogen ion and then diffuse into the next cell.

Once in a cytoplasm again, auxin loses the H^+ ion and becomes negatively charged.

It can then again move to the next cell, and therefore redistribute to the shadier side of the plant.

Once in the shadier parts of the plant, auxin binds its receptor and activates genes responsible for secretion of more H^+ ions into the cellulos cell wall.

With the extra H^+ ions in the cell wall, the acidity increases, and the cellulose loosens up, allowing the plant to elongate, which leads to the general movement of the plant towards the light. This is because when the cells elongate, they cause a bending of the plant towards the opposite side.



Phalaris canariensis: cotyledons after exposure in a box open on one side in front of a south-west window during 8 h. Curvature towards the light accurately traced. The short horizontal lines show the level of the ground.

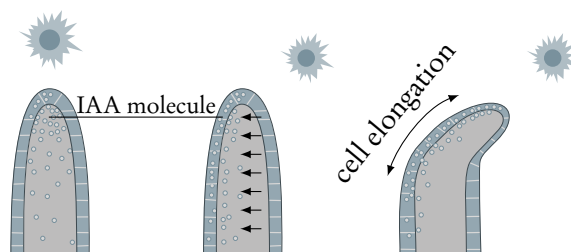


Figure 9.6: Auxin.

In the roots, however, auxin inhibits cell elongation.

9.4.3 Germination

In order for the embryo shoot to emerge from the seed (recall section 9.3), the seed needs to be exposed to optimal conditions

Three factors play a major role:

Water to rehydrate the plant seed

Oxygen to start up cell respiration

Warmth to provide optimal temperature for enzyme activity

You can design an experiment where you provide your seed with enough water, oxygen and heat. In order to determine the effect of each factor, you should keep the other two constant, and vary the third. Here is an example.

Example.

Effect of water

Take 2 sealed flasks and place them at room temperature.

Make sure that the flasks are large enough to provide enough oxygen for several days when sealed.

Before sealing, add a moist cotton wool at the bottom of one, and a dry cotton wool at the bottom of the other.

Place the seeds on the cotton wool, and seal the flasks.

Examine the germination over a period of a few days.

9.4.4 Micropropagation



Micropropagation is a term used to describe propagation of plants (growth) from a single small piece of plant tissue.

First, a small piece of plant tissue is taken from the shoot tip of a plant.

The piece of tissue is sterilised and placed in a sealed flask containing nutrient rich agar gel with abundant auxin.

The plant is allowed to grow into a shapeless lump of tissue called callus which can be split for extended growth.

A piece of callus can be transferred to auxin poor agar, containing other hormones needed for shoot and root development.

Once the plantlet develops, it can be transferred to soil to continue its growth into a proper plant.

Advantages of micropropagation

- It speeds up the process of plant propagation by using only small pieces of tissue and supplying them with appropriate hormones
- The shoot tips usually are virus free, even though the rest of the plant can be infected, so virus-free strains can be produced
- The costs of plant production is decreased, and plants needn't be taken out of their natural habitat

10.1. Antibody production and vaccination **194**

- Antibody production
- Production of monoclonal antibodies
- Allergies
- Pathogen specificity for host infection

10.2. Movement **198**

- Bones and exoskeletons
- Synovial joints
- Skeletal muscle structure
- Muscle contraction mechanism

10.3. Kidneys and osmoregulation **203**

- Kidney
- Ultrafiltration
- Selective reabsorption
- Salt/water regulation
- Osmoregulation by ADH
- Comparing the filtrate and urine composition
- Comparing the content of renal artery and vein
- Kidney failure
- Osmoregulation in animals
- Malpighian tubule

10.4. Reproduction **212**

- Spermatogenesis
- Oogenesis
- Comparing spermatogenesis and oogenesis
- Fertilization
- Embryo development
- Pregnancy hormones
- Placenta

10.1 Antibody production and vaccination

10.1.1 Antibody production

Recall that the specific response to pathogens was mounted by lymphocytes via the antibody production. Lymphocytes are divided into T and B cells; T cells are the so called helper cells that activate B cells, which in turn produce the specific antibodies.

When pathogens are encountered, phagocytes engulf and break down pathogens. These cells will engulf and break down anything recognized as “non-self.”

After engulfment, these “antigen-presenting cells” will display fragments of the broken down pathogen on their surface, and migrate to the lymph nodes to alert the immune system of a threat.

T helper cells contain T cell receptor molecules on their surface that bind specific antigens. These cells will then get activated by the phagocyte’s warning and release cytokines to activate particular B lymphocytes.

The activated B lymphocytes are capable of producing the specific antibodies for the recognized antigen that was presented by the phagocyte.

These activated B lymphocytes will rapidly divide into high numbers of short-lived plasma cells that release large numbers of antibodies into the blood stream to target the pathogens for destruction.

A small proportion of the activated B and T cells will become memory cells, which provide long lasting immunity to the pathogen. Therefore, in the case of a subsequent infection, these cells will quickly react to produce large numbers of antibodies much faster.

Antibodies are crucial to the immune system as they prevent pathogens from invading healthy cells, target pathogens for removal, and initiate certain destruction pathways.

In order to have B lymphocytes capable of releasing antibodies for any antigen, the body needs to produce random antibodies that could potentially fit to any pathogen that it may encounter

- Antibodies are made of 4 polypeptide chains, two heavy and two light, that are joined via disulphide bonds to make a Y shape (figure 10.1).
- The heavy chains are constant regions of the antibodies, meanwhile the light chains are termed “variable regions.”
- Variables regions get their name because they are different for all antibodies, made in a random manner by each B lymphocyte

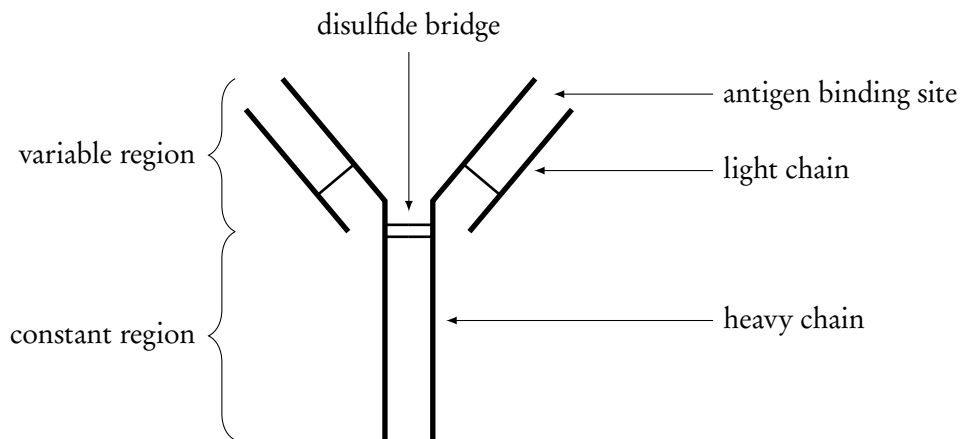


Figure 10.1: Antibody structure

- In this way, the body has all sorts of antibodies with different variable regions for high specificity to any antigen in the body

The functions of the antibodies are the following:

- Binding the antigens on pathogens makes these pathogens more visible to immune cells.
- Binding the virus antigens can prevent virus entry into the cell.
- The toxins are produced by pathogens, are also antigens on their own, and can be neutralized by antibody binding.
- By binding many antigens on a pathogen's surface, the antibodies can create holes in the cell and kill it.
- Antibodies can cause the bound pathogens to stick to each other, making them less mobile and disabling them from entering the host cells.



Vaccines are a modified, weakened form of a pathogen that contain antigens from that pathogen.

Once the vaccine is injected, the antigens stimulate a specific immune response called a primary response. These antigens stimulate macrophages T-cells which in turn stimulates cloning of B-cells including development of memory (B-)cells that produce specific antibodies.

Upon subsequent exposure the productions of antibodies is much faster initiating what is called a secondary response. Thus the person is said to have immunity.

Sometimes, vaccines are given a few times, since the first vaccination only produces a mild response with few memory cells

The second, booster vaccination, triggers the memory cells to launch a full response, and to produce more memory cells, this is an example of active immunity.

Example.**Jenner's smallpox experiment**

Jenner performed a controversial experiment where he infected an 8 year old child with cowpox which is a less severe, but very similar version of the smallpox virus.

The child fought the cowpox infection, and upon exposure to smallpox, the child didn't get infected.

He eventually tested this on other people and himself, but his methods would today be considered unethical since he had tested his idea directly on humans, without prior animal and safety tests.

10.1.2 Production of monoclonal antibodies

Based on the principles of antibody production in the body, an ingenious method was developed to produce specific antibodies outside of the human body.

The idea is to inject a type of antigen into an animal, and collect the B plasma cells that the animal has produced.

These cells can then be fused with tumour cells and produce hybridomas, which divide continuously, therefore producing large quantities of antibodies of that antigen.

The antibodies that are secreted can be purified and used for diagnostics or treatment.

One use of monoclonal antibodies is in pregnancy tests. This uses the antibody against HCG, the hormone produced during pregnancy and excreted through urine.

10.1.3 Allergies



Allergies are immune responses to harmless substances, such as pollen, food, etc.

Allergies occur when immune cells get overstimulated by these substances and produce large quantities of histamines.



Histamine is a molecule that causes vasodilation in infected areas in order to promote white blood cells researching the site of inflammation.

Basophils and mast cells produce histamines, which is normally a useful substance, but upon over-activation of these cells, it causes symptoms associated with allergies, such as rash, sneezing, itching and in more serious scenarios, anaphylaxis, a dangerous swelling in the air canals. Anti-histamine drugs are used to alleviate symptoms of allergies.

10.1.4 Pathogen specificity for host infection

Species specificity of pathogens means that not all pathogens can infect all species, and this can be dependent on their surface molecules, modes of infection, and immune barriers of the host.

For example, Measles, Polio and Syphilis occur only in humans, while tuberculosis can infect other animals, such as cattle which can transmit it to humans.

The same case occurs in rabies, which can be transferred from infected dogs onto humans, through saliva.



Zoonosis is when a disease can be transferred from humans to other animals.

10.2 Movement

10.2.1 Bones and exoskeletons

While many other structures are involved in movement, the attachment of muscles to bone and/or exoskeleton is essential to control and initiate movements. This is because bones and exoskeletons:

provide a firm anchorage for muscles,

act as levers,

have the capacity to change the size or direction of the forces generated by muscles.

The movement of the body requires muscles to work in antagonistic pairs, meaning that when one contracts the other relaxes. Muscles attached to static bones must then contract in order to bring about one movement while the antagonistic muscle contracts to bring about the opposite movement (for example an arm flexion and extension).

10.2.2 Synovial joints

Joints are composed of several structures. Their names and functions are:

Bones: provide anchorage and act as levers.

Ligaments: connect bone to bone, restricting movement at joints and helping to prevent dislocation.

Tendons: cords of dense connective tissue that attach muscles to bone.

Muscles: provide the force needed for muscle contraction. In most cases, this force comes from the action of antagonistic muscle pairs (where one muscle contracts and the other relaxes in unison to elicit movement).

Nerves: stimulate muscles to contract at a precise time and extent, so that movement is coordinated.

Try it now! Your elbow can only move sideways and back and forth, whereas the hip allows for broader circular movements as well!

In order to visualize and understand the structure and importance of the structures present in joints, Figure 10.2 is a labelled diagram of the human elbow, as well as a brief description of the components shown in the diagram.

Cartilage: a layer of smooth and tough tissue that covers the ends of the bones where they meet to reduce friction.

Synovial fluid: lubricates the joint to reduce friction and provides nutrients to the cells of the cartilage.

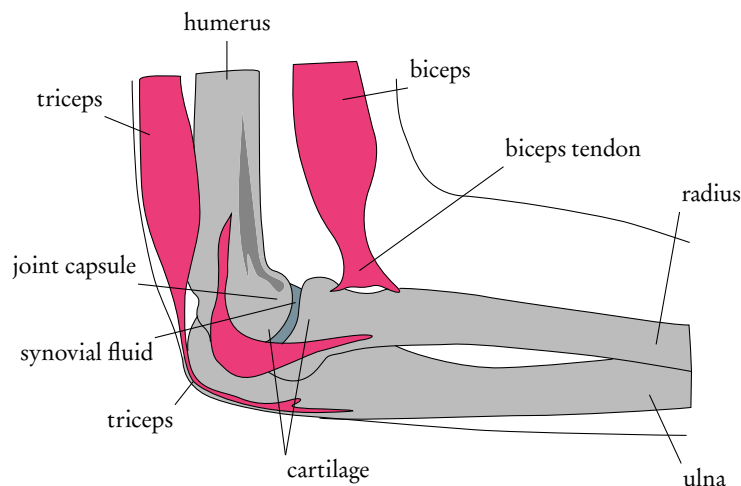


Figure 10.2: Elbow joint.

Skill: Annotation of a diagram of a human elbow.

Joint capsule: surrounds the joint, encapsulates the synovial cavity and unites the connecting bones (it “seals” the joint).

Humerus: acts as a lever that allows anchorage of the muscles of the elbow.

Radius: bone that transmits forces from the biceps through the forearm.

Ulna: bone that transmits forces from the triceps through the forearm.

Biceps: the flexor muscle, used to bend the arm at the elbow.

Triceps: the extensor muscle, used to straighten the arm. These two muscles act as antagonistic pairs!



Synovial joints are capsules that surround the articulating surfaces of two bones (where two bones meet)

TIP: to remember which bone is which, place your arm flat on the desk and do a thumbs up, the radius is the top bone, and the ulna rests directly against the surface of the desk.

There are various types of synovial joints, each with characteristic patterns of movement:

- Ball-and-socket joint (like the hip), which allows movement in three planes.
- Hinge joint (like the knee and elbow), which allows movement in two planes.

This means that synovial joints allow certain movements, but not all.

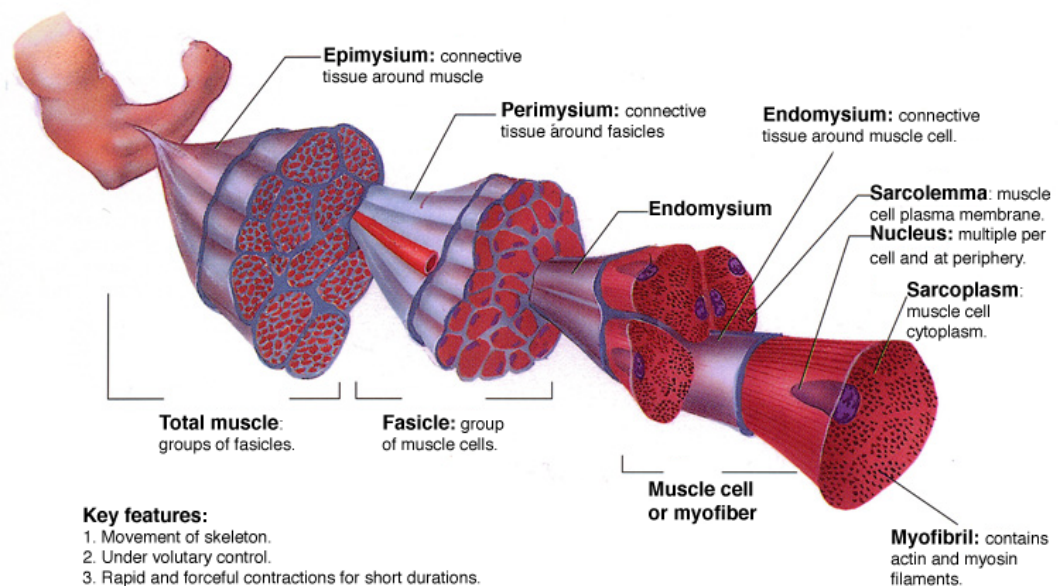
10.2.3 Skeletal muscle structure



Skeletal Muscle is a type of striated muscle that is under the voluntary control of the somatic nervous system.

Skeletal muscle consists of large multinucleate cells called muscle fibres, which have an elongated, cylindrical shape. These cells contain multiple nuclei found within the fibres' sarcoplasm (very much like the cytoplasm of a cell), which is surrounded by the sarcolemma (its plasma membrane). Muscle fibres also contain a sarcoplasmic reticulum, from which calcium is released upon nerve stimulation.

Skeletal Muscle



Within each muscle fibre, there are smaller cylindrical structures called myofibrils, which consist of repeating units called sarcomeres (you can see the structure of a sarcomere in Figure 10.3).

Sarcomeres contain filaments of contractile proteins actin and myosin, both very important in the process of muscle contraction.

Skill: Drawing labelled diagrams of the structure of a sarcomere.

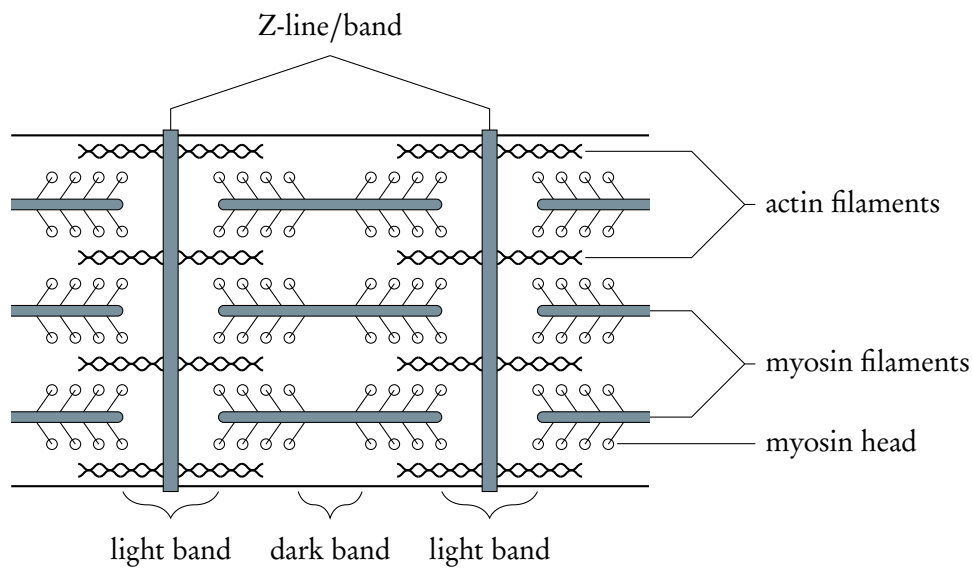


Figure 10.3: Sarcomere.

Z-line/band: structure to which narrow actin filaments are attached, when actin slides over thick myosin filaments, the Z-bands move closer to each other.

Light band: the region of the sarcomere where only the thin actin filaments are present.

Dark band: the region of the sarcomere where thick myosin filaments are present (alone or in overlap with actin).

Actin filaments: thin fibrous proteins that contain myosin binding sites.

Myosin filaments: thicker fibrous proteins that have “heads” that are capable of attaching to the neighbouring actin filaments.

10.2.4 Muscle contraction mechanism

The mechanism of muscle contraction is explained by the “sliding filament theory”, which states that muscles contract when actin filaments slide over myosin filaments.

Muscle contraction is guided by three main players: calcium ions, troponin, and tropomyosin. A complex of troponin and tropomyosin covers the myosin binding sites in actin filaments when the muscle fiber is at rest. Troponin has binding sites for calcium that, when occupied, cause it to change configuration of the complex and uncover the myosin binding sites. Recall that when a nerve impulse reaches the muscle to cause a contraction, calcium is released from the sarcoplasmic reticulum, initiating the action.

The sliding filament theory can be divided into 6 main steps:

1. The arrival of an action potential triggers the release of Ca^{2+} ions from the sarcoplasmic reticulum
2. Calcium ions bind to troponin and reconfigure the complex, exposing the binding sites for the myosin heads.

3. Myosin head binds to the binding sites, creating a cross-bridge where the myosin head is at a high energy configuration (fig 6.15 top) as it has both ADP and phosphate bound to it.
4. Myosin head pivots and releases the ADP and phosphate (fig 6.15 left). This results in a lower energy configuration and pulling of the actin filament toward the M line.
5. A new ATP binds to the myosin head, causing the cross-bridge to break (fig 6.15 bottom).
6. ATP hydrolysis to ADP and phosphate causes the myosin head to cock back (fig 6.15 right). From here, steps 3-6 repeat until Ca^{2+} is released from troponin.

We can analyze electron micrographs of relaxed and contracted muscle. As shown in the Figure 10.5, when skeletal muscle is contracted the light bands become much smaller, as more actin is sliding over myosin filaments. Also, the Z-bands are closer together, as actin moves closer towards the centre of the sarcomere.

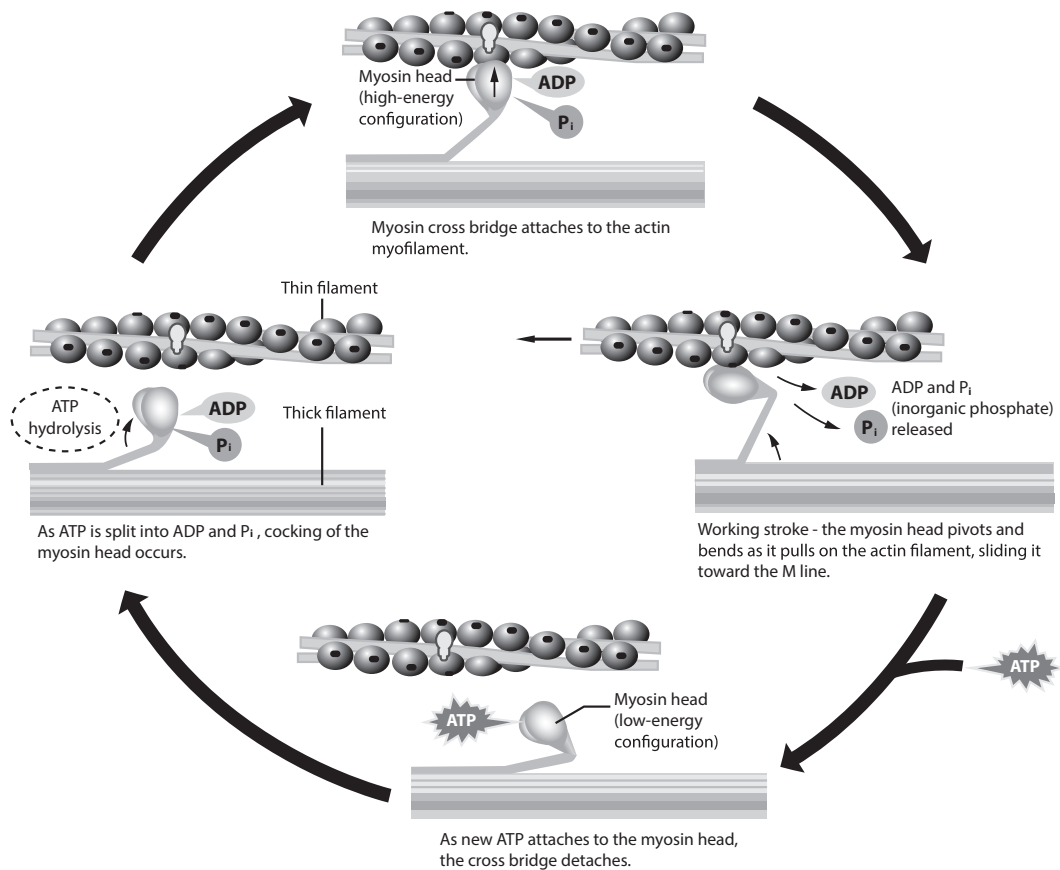


Figure 10.4: Muscle contraction mechanism.

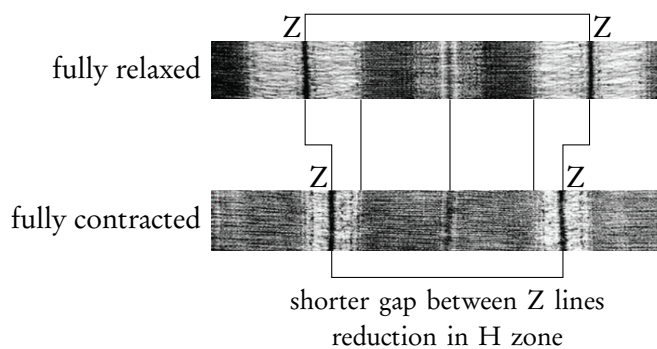


Figure 10.5: Muscle contraction and relaxation.

Skill: Analysis of electron micrographs to find the state of contraction of muscle fibres.

10.3 Kidneys and osmoregulation



Excretion is the process of removal of potentially toxic waste products of metabolic pathways.

Nitrogenous wastes are compounds produced through metabolic pathways and excreted by animals.

Nitrogenous wastes include three groups of compounds:

Ammonia excreted by freshwater fish and amphibian larvae;

- ammonia is a very toxic compound which can only be excreted as a very dilute solution with a lot of water;
- ammonia is excreted by animals that live in water rich habitats, such as fish, since the water is abundant there.

Urea excreted by terrestrial and marine mammals, marine fish and adult amphibians;

- urea is not as toxic, and can be excreted with smaller quantities of water;
- conversion of ammonia to urea requires energy, but animals that do not have an abundant supply of water need this conversion to prevent excessive water loss.

Uric acid excreted by birds and insects;

- uric acid is not toxic even at its highest concentrations, and is excreted in a form of a paste;

Metabolic pathways, as you will recall, are chemical pathways and cycles which cells use to breakdown or build-up molecules.

- conversion of ammonia to uric acid requires even more energy, but birds take advantage of that, since concentrating urea into uric acid, reduces water mass that would normally be taken up by water containing urea.

Recall that metabolic pathways are composed of chemical reactions that not only produce waste products, but often consume or produce water. This causes water levels to be always changing within an organism, which affects the osmotic properties of the cells in their bodies. According to how they manage their internal osmotic concentrations, organisms can be classified as osmoregulators and osmoconformers.



Osmoregulators keep their body's osmolarity constant, regardless of their environment.

Osmoconformers keep their body's osmolarity equal to the osmolarity of the environment.

Most osmoconformers are marine organisms, where maintaining a body osmolarity equal to the aqueous environment ensures minimal influx and efflux of water from their cells, and requires minimal energy. Osmoregulators, in contrast, spend a lot of energy to tightly regulate their optimal internal conditions and not be heavily affected by changes in the environment. Humans and most terrestrial animals are osmoregulators.

10.3.1 Kidney



Nephrons are the functional units of the kidney that spans the regions of the cortex and the medulla.

Each nephron spans the cortex with its glomerulus and proximal and distal tubules, and spans the medulla with the loop of Henle and collecting duct.

Glomerulus is a cluster of capillaries that are contained at the Bowman's Capsule.

Bowman's Capsule is associated with the glomerulus, and it is the first site of filtration of the blood.

Proximal convoluted tubule is responsible for the selective reabsorption of nutrients from the filtrate.

Loop of Henle is the site of salt and water reabsorption into medulla.

Distal convoluted tubule further regulates solute concentration of blood (and its pH) through selective reabsorption.

Collecting duct is the site of final water reabsorption or excretion by the nephron.

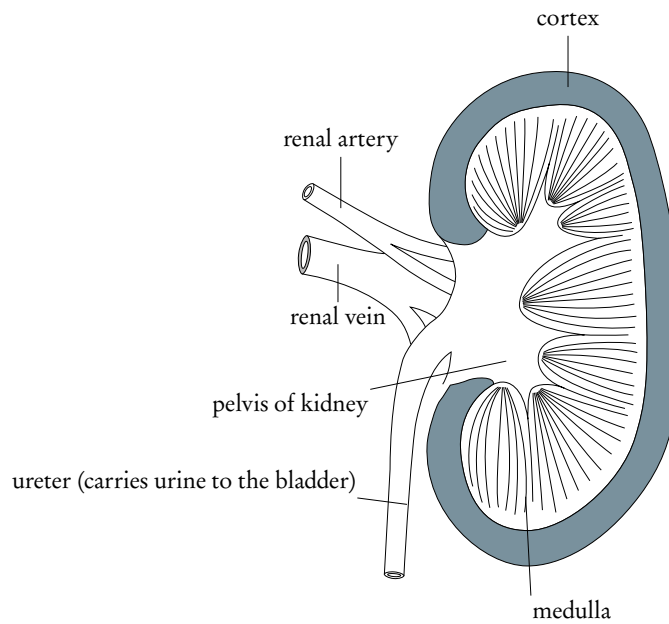


Figure 10.6: Kidney.

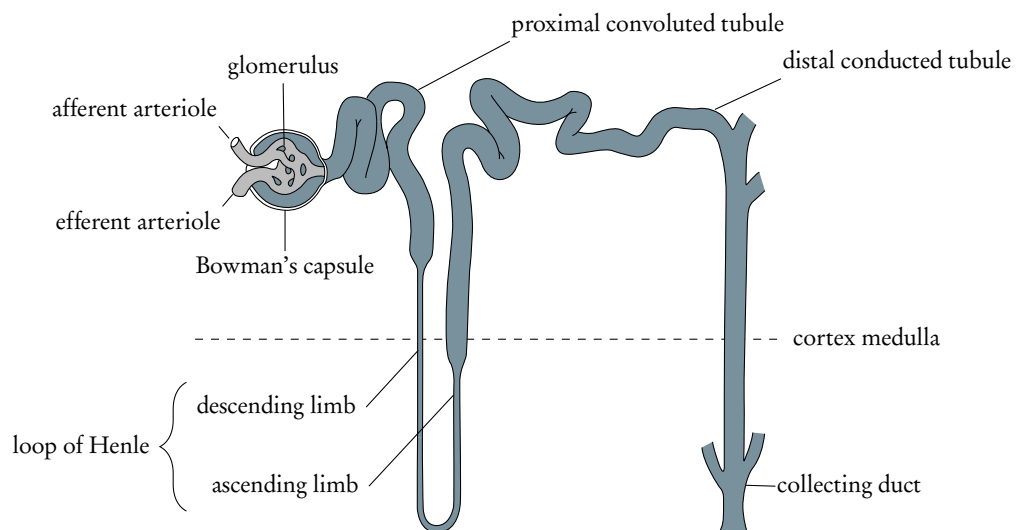


Figure 10.7: Nephron.

10.3.2 Ultrafiltration

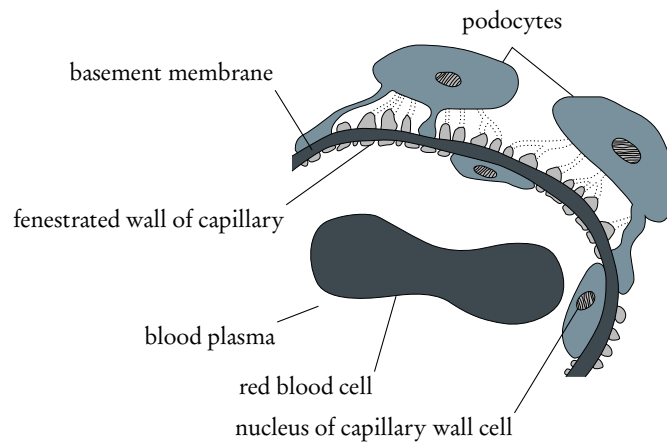


Figure 10.8: Ultrafiltration.

Ultrafiltration is the non-selective filtration of the blood and it occurs at the Bowman's capsule of the nephron. Here, the filtration process is driven by the pressure gradient at the glomerulus, which arises from the differences between the afferent and efferent arterioles. The afferent arteriole brings blood into the glomerulus and it has a much larger diameter than the efferent arteriole, which takes the filtered blood away from the glomerulus. Due to the high pressure created between the two arterioles in the glomerulus, a lot of nutrient rich fluid escapes from the blood into the Bowman's capsule.

In addition, in order for nutrients to effectively go from the capillaries to the nephron, certain structures are necessary:

- The capillaries at the glomerulus are (bold) fenestrated, meaning that they have pores from which blood can freely exit the vessels.
- The Bowman's Capsule is lined with cells called podocytes. Podocytes wrap capillaries with their foot processes, and the gaps formed in between these processes are called filtration slits. These slits allow fluid to freely enter the nephron.
- The basement membrane is a mesh-like structure in between the glomerulus and the podocytes. It is size-selective, meaning that it is this structure that determines which molecules will move from the capillaries into the nephron.

Overall, ultrafiltration leads to a filtrate containing small blood soluble molecules, such as glucose, salts and nitrogenous wastes. But proteins, red blood cells, and other blood cells, remain in the capillary.

10.3.3 Selective reabsorption

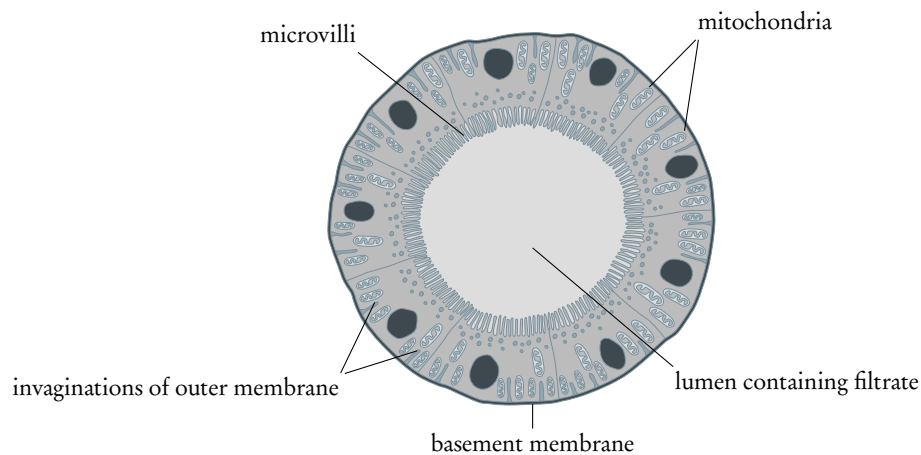


Figure 10.9: Selective reabsorption.

Selective reabsorption occurs in the proximal convoluted tubule (PCT). The function of the PCT is to reabsorb all the molecules from the filtrate that should not be excreted out of the body (like glucose). The PCT is made of a single layer of cells with microvilli pointing toward the lumen. The cells contain many mitochondria to provide enough energy for the active transport of molecules from the filtrate into the cells. 80 percent of the ions (such as sodium and potassium) from what initially enters the nephron at ultrafiltration are reabsorbed into the bloodstream at the PCT.

Since the cells have a relatively small volume compared to the volume of the entire tubule, the solute concentration in them is much higher, so the water from the tubule follows into the cells by osmosis.

Additional reabsorption occurs at the distal convoluted tubule, mainly dictated by hormonal changes.

10.3.4 Salt/water regulation



Isotonic is a term describing equal solute concentrations inside and outside of the body.

Hypotonic is a term describing a higher solute concentration inside the body (compared to the environment).

Hypertonic is a term describing lower solute concentration inside the body (compared to the environment).

Overhydration results from too much water entering an organisms, and can lead to behavioural changes, confusion, muscle cramps, nausea, coma and death.

Dehydration results from an insufficient water supply to the body, leading to thirst, excretion of concentrated urine, tiredness, low blood pressure, brain damage and death.

If you recall the structure of a single nephron, you will notice that the loop of Henle runs mostly in the medulla, unlike glomerulus and convoluted tubules which are entirely situated in the cortex.

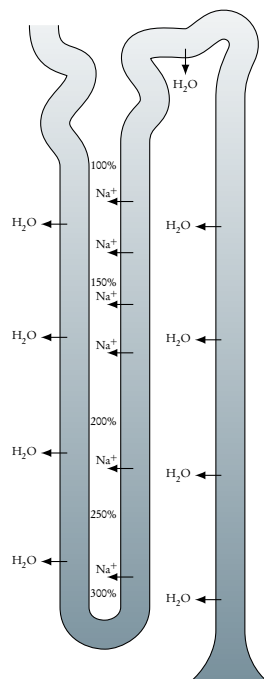


Figure 10.10:
Loop of Henle.

The salt and water absorption mainly occurs in the loop of Henle, which is a U shaped structure with a descending and an ascending limb.

As the filtrate flows through the tubule, the salts are being pumped out into the medulla to create a high salt concentration which then pulls the water out of the filtrate through osmosis. Descending limb is permeable to water, and impermeable to sodium.

Ascending limb is permeable to sodium, but not to water. The filtrate flows through the descending limb where the water is pulled out into the medulla by osmosis. But how? Since there is always a continuous stream through the entire nephron, the filtrate that is in the ascending limb is losing its sodium to the medulla, which makes the entire medullary area highly concentrated.

This reflects to the descending limb which is permeable to water, so the salt that is excreted in the ascending part causes the water in the descending part to enter the medulla.

Looking at the picture of the Loop, you will notice that as the filtrate flows down the loop, it becomes more and more concentrated since water is being lost, but as it flows up, the ions are being pumped out, so the concentration again decreases.

The concentration of medulla is then referred to as hypertonic.

10.3.5 Osmoregulation by ADH

Osmoregulation occurs in the last part of the nephron, the collecting duct. When the body senses that the blood does not contain enough water, the pituitary gland in the brain secretes a hormone called anti-diuretic hormone (ADH).

ADH travels in the blood to the collecting duct where it stimulates the cells to produce more water channels called aquaporins (water pores).

Due to the high osmolarity at the medulla (previously achieved by the loop of henle), the water pores will allow for a big volume of water to exit the duct and enter the tissue.

This will lead to a small volume of highly concentrated (more yellow) urine to be produced.

If the body doesn't need extra water retention, ADH is not produced, and the collecting duct cells allow little water to leave into the medulla. This results in a large volume of diluted urine.

10.3.6 Comparing the filtrate and urine composition

Now that you know what part of the kidney does what, and what substances can be found in which part, it won't be difficult to compare the composition of filtrate and urine at different stages of the nephron.

Table 10.1

	Blood in glomerulus	Filtrate in glomerulus	Filtrate in loop of Henle	Filtrate in distal convoluted tubule	Urine with ADH	Urine without ADH
Proteins	Abundant	None	None	None	None	None
Glucose	Abundant	Abundant	None	None	None	None
Urea	Some	Some	Concentration increasing	Concentration increasing further	Very concentrated	Less concentrated

Notice that the proteins cannot pass into the filtrate, glucose must be entirely absorbed before the loop of Henle, and the concentration of urea increases through the filtrate, but varies with amount of water being retained or excreted.

10.3.7 Comparing the content of renal artery and vein

Again, based on the known functions of the nephron, it should be simple to explain the differences in composition of arterial and venous renal blood.

Oxygen Artery has a higher concentration compared to the vein, since the kidney uses oxygen for aerobic respiration.

CO₂ Lower in the artery, compared to the vein, since the vein produces more CO₂ due to aerobic respiration.

Glucose Only slightly higher in the artery, since some of the glucose is used up by the kidney for respiration.

Urea High urea concentration in the artery, and lower in the vein, due to excretion.

Plasma proteins The concentrations are equal, since no proteins can pass into the filtrate.

Na/Cl ions Concentrations can vary in the artery, but since the function of the kidney is to regulate salt concentration, the venous blood always contains normal amounts of salts.

These are the normal conditions in a healthy person, but some diseases might alter the composition of blood and urine.

- Blood cells in urine can be a sign of an infection or cancer.
- Glucose in urine is a sign of diabetes, since only extremely high levels of blood glucose result in insufficient reabsorption by the proximal tubule.
- Proteins in urine are a sign of glomerular damage, since they are too large to pass a healthy membrane.

10.3.8 Kidney failure

If the kidney cannot filter the blood, the body will fill up with toxins that can lead to death. Kidney failure can be treated in two ways, by *hemodialysis* or *kidney transplant*.

An alternative to filtration in these cases is hemodialysis. In this process, the blood from the patient is passed through a machine that imitates the nephron. The machine contains dialysate fluid which is composed of exactly the right amounts of salts, glucose and ions to filter blood. The blood passes through tubing with a semi-permeable membrane which is in contact with dialysate. No urea is present, so that all urea can leave the patient's blood and end up in the dialysate.

Glucose is set at a concentration where the body will not lose any of the nutrients to the dialysate.

High Ca²⁺ and low K⁺ concentrations ensure that calcium is absorbed into the blood and potassium excreted.

Salts are set at the concentration to ensure the body is properly hydrated.

Hydrogencarbonate ions are added to buffer the acidity of the blood.

Dialysis has to be performed 3 times a week for several hours, but is not a permanent solution to people with kidney failure.

Kidney transplant is a long term solution that requires a donor (could be recently deceased or a living match) to donate one kidney to the sick patient.

The most important aspect is that the donor and receiver are a good match in blood group, in order to avoid immune rejection by the host organism.

10.3.9 Osmoregulation in animals

Recall that the loop of Henle is the main water and salt absorbing area of the nephron. Also recall that the loop of Henle extends into the medulla.

Research in animals that live in different habitats has shown that animals that live in more water deprived habitats, have thicker medulla than the ones living in aquatic habitats.

The reason for this is the fact that the thicker the medulla, the longer the loop of Henle can be, and therefore, the more efficient water retention and conservation. Therefore, the length of the loop of Henle is said to be correlated with the need of water conservation.

10.3.10 Malpighian tubule

Insects do not have a blood-based system, but a hemolymph system which employs a long vessel from the back to the front of the body.

The vessel is used to transport hemolymph, is a fluid equivalent to blood, along the body and release it into various parts of the tissue.

The hemolymph then has to diffuse through the tissue until reaching the main vessel again for a new cycle.

In order to collect the waste from the hemolymph, long projections called malpighian tubules, enter the tissue in the middle part of the body, and absorb the wastes from the hemolymph that has been pumped into the tissue.

The wastes have to be returned to the gut and excreted, so the tubules contain special ion channels that cause the ions to enter the tubule.

The water follows the ions leading to the flow of wastes back to the gut.

In the gut, the wastes travel to the rectum where the ions are again pumped out into the hemolymph.

The water follows, leading to water and ions conservation, and waste excretion.

10.4 Reproduction



Gamete is a haploid cell that fuses with another haploid cell during fertilization (in organisms that have sexual reproduction)

Spermatogenesis is the production of male gametes in the testes.

Oogenesis is the production of female gametes in the ovaries.

10.4.1 Spermatogenesis

- The first stage of sperm production requires growth and divisions of spermatogonium by *mitosis*. This division produces a primary spermatocyte and another spermatogonium (a kind of stem cell).
- The primary spermatocyte grows and then enters meiosis. After completion of meiosis I, it forms two secondary spermatocytes.
- The secondary spermatocytes undergo meiosis II to form four spermatids.
- Spermatids (haploid) undergo a process of differentiation and maturation in which they become sperm cells, which are motile and have the required organelles.

10.4.2 Oogenesis

- This process begins during fetal development.
- The first stage requires growth and divisions of primordial cells (a kind of stem cell) by *mitosis*. This division produces primary oocytes and more primordial cells.
- Primary oocytes begin meiosis, but are arrested at prophase I as granulosa cells begin surrounding the oocytes to form follicles. These follicles remain arrested in prophase I until women begin puberty.
- Each month with the menstrual cycle, FSH will trigger meiotic division of some of the follicles.
- The first division leads to two daughter cells of unequal size, as one egg (the secondary oocyte) has most of the cytoplasm and organelles, the other one is the polar body. Polar body stays by the granulosa cells until it degenerates.
- The secondary oocyte undergoes meiosis II, but is arrested at metaphase II and is released from the ovary to the fallopian tube to await from fertilization. The secondary oocyte is surrounded by the follicle cells that nourish and protect it.

- If fertilization occurs, a chemical reaction causes completion of meiosis II, leading to the formation of an ovum and a second polar body that will degenerate. The cell is an ovum for the moment before it fuses its genetic material with the sperm and then becomes a zygote.

10.4.3 Comparing spermatogenesis and oogenesis

Table 10.2: Comparing spermatogenesis and oogenesis.

	Spermatogenesis	Oogenesis
<i>Number of gametes produced</i>	many / millions per day	one per month/menstrual cycle/28 days / about 400 eggs per life time
<i>Products of meiosis</i>	four / equal division of the cytoplasm / no polar bodies	one / unequal division of the cytoplasm / polar bodies
<i>Start of process</i>	at puberty	begins during fetal development
<i>Duration of production</i>	throughout adult life	ends at menopause
<i>Timing of release</i>	produced continuously / released during ejaculation	released at ovulation / in the middle of the menstrual cycle
	both spermatogenesis and oogenesis involve meiosis and mitosis	
	both produce haploid cells/nuclei	
<i>Both occur in gonads</i>	occurs in testes	occurs in ovaries

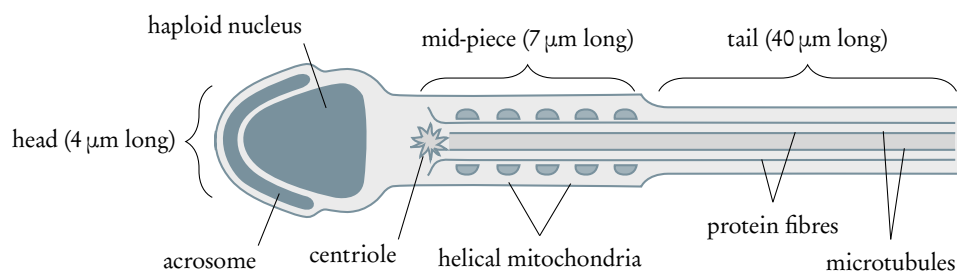


Figure 10.11: Sperm cell.



Acrosome is found in the head of the sperm, contains enzymes to digest zona pellucida of the egg.

Nucleus contains haploid number of chromosomes (23).

Tail is used for the movement of the sperm cell towards the egg in the oviducts, with the help of microtubules.

Mitochondria provide the energy for the swimming.

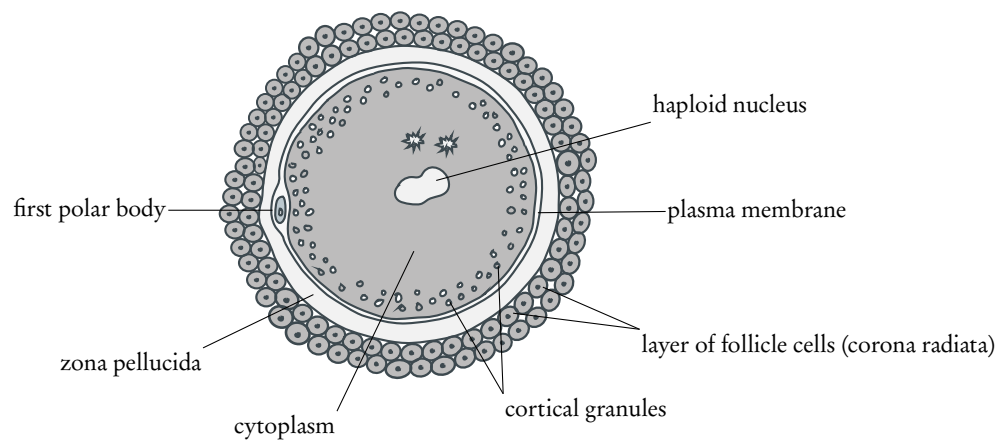


Figure 10.12: Egg cell.



Nucleus contains haploid number of chromosomes.

Cytoplasm contains abundant nutrients to feed the early embryo. It is the cytoplasm of the egg cell that ultimately harbours the zygote

Cortical granules are used to prevent polyspermy by hardening the zona pellucida after fertilization.

Zona pellucida is a protective layer that limits sperm entry.

10.4.4 Fertilization

Fertilization can be both internal or external, and it relates to the environment where the organism lives.

Internal fertilization occurs when the male passes the sperm directly into the female body, to join the egg cells (mammals, some birds, pythons and reptiles).

External fertilization occurs when the female lays eggs in the environment, and the male covers the eggs with sperm, outside the female body (salmon, fish, frogs).

This is possible in aqueous or moist environments where the sperm can freely move toward the egg.

Stages of fertilization

Fertilization in humans can be divided into three main stages: capacitation, acrosome reaction, and cortical reaction.

1. Capacitation: biochemical changes post ejaculation to improve sperm motility.
 - Uterine chemicals dissolve a cholesterol coat around the sperm, which improves motility and destabilizes the acrosome cap which aids in acrosomal reaction.
2. Acrosome Reaction: release of hydrolytic enzymes that soften the zona pellucida.
 - When the sperm reaches the egg, the reaction is responsible for the breaking of the zona pellucida
 - Acrosome vesicle fuses with jelly coat and releases hydrolytic enzymes so the sperm can reach the egg membrane's dock proteins.
 - Membrane of egg and sperm fuse and sperm nucleus enters the egg.
3. Cortical reaction: hardening of the zona pellucida.
 - Occurs when sperm has penetrated the egg.
 - Cortical granules in the egg cytoplasm release enzymes into the zona pellucida to destroy dock proteins and harden the glycoprotein matrix of the jelly coat.
 - Ensures that no other sperm fertilizes the egg, therefore prevents polyspermy.



Polyspermy is when several sperm cells fertilize one egg. Polyspermy must be avoided because the fusion of two haploid nuclei leads to formation of a diploid cell, but a fusion of three (or more) haploid nuclei leads to a triploid (or more) cell that cannot survive.

10.4.5 Embryo development

Fertilization usually takes place in the oviducts.

After fertilization, an influx of calcium promotes the completion of meiosis II of the egg, resulting on the zygote (egg and sperm genetic material). The zygote begins quick mitotic divisions, making two-, four-, eight- (etc.) cell embryo.

After many cycles of division, the developing ball of cells becomes a blastocyst, which has an inner cell mass surrounded by an outer layer of cells (fig X). This early embryo will eventually implant in the uterine wall to develop into the placenta (outer cell layer) and embryo (inner cell layer)

10.4.6 Pregnancy hormones

Once the embryo has implanted into the uterine lining and started developing, it starts secreting hCG hormone which leads to continuous production of progesterone by the ovaries.

Progesterone maintains the thickness of the uterine lining.

After 12 weeks, the ovaries cannot keep up with the progesterone production, but by then, the placenta has already developed so it takes over progesterone and oestrogen production.

Progesterone also prevents contractions of the uterus, but after 9 months, the levels start to fall.

Drop in progesterone leads to secretion of oxytocin.

Binding of oxytocin to its receptors causes the uterus to contract, and the contractions cause more oxytocin to be produced (positive feedback).

While the contraction continue and become more frequent, the cervix relaxes and the amniotic sac breaks.

Once the baby is delivered, the umbilical cord is cut.

The contractions continue until the placenta is delivered as well.

10.4.7 Placenta

The placenta provides a connection between the mother and child's circulations.

When the embryo is 8 weeks old, it is termed fetus and it develops the placenta and umbilical cord.

The structure of the placenta is the following:

- Myometrium and endometrium are the outer and inner walls of the uterus, respectively.
- Placenta originates from the fetus, and is embedded in the endometrium.
- The placenta contains placental villi which are projections containing the fetal capillaries.
- The space between the villi is called inter-villous space, and contains pools of the mother's blood.
- The blood of the mother is always separated from the child's blood by the chorion.

The umbilical cord contains two main fetal arteries and a fetal vein, so that the arteries can take deoxygenated blood to the mother, and the vein can take the oxygenated blood to the baby (recall that arteries take the blood away from the heart, and veins bring it to the heart).

As mentioned, the child's blood is brought in capillaries to the villi, where the blood makes small pools from the other side of the chorion.

The chorion is the barrier between the two circulations and it contains many microvilli and mitochondria to facilitate active transport.

The chorion also produces progesterone and oestrogen.

To minimize the transport distance, the capillaries, as well as the chorion, are one cell thick, so the exchange of materials can happen quickly.

The child receives oxygen, nutrients, antibodies and minerals from the mother, and releases carbon dioxide, urea and hormones to the mother's circulation.

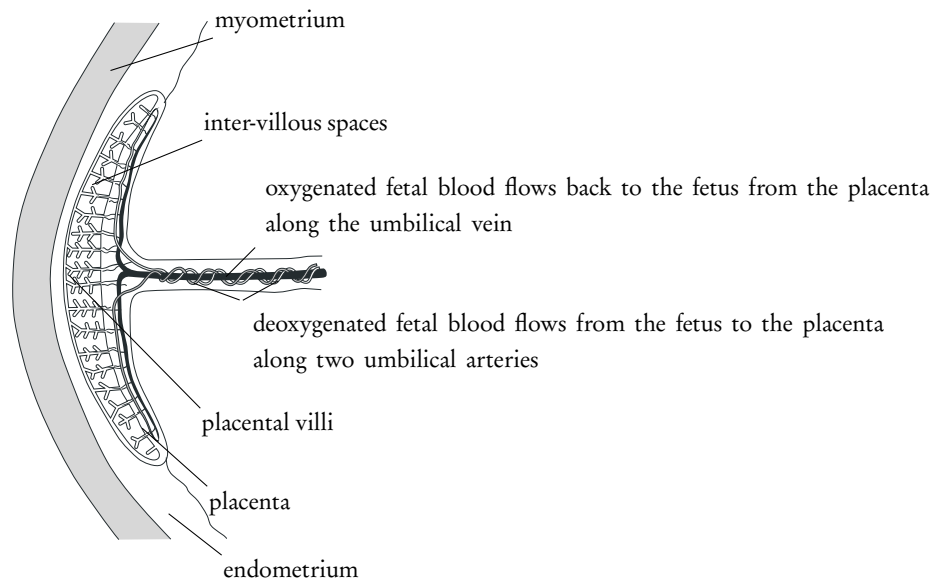


Figure 10.13: Placenta.

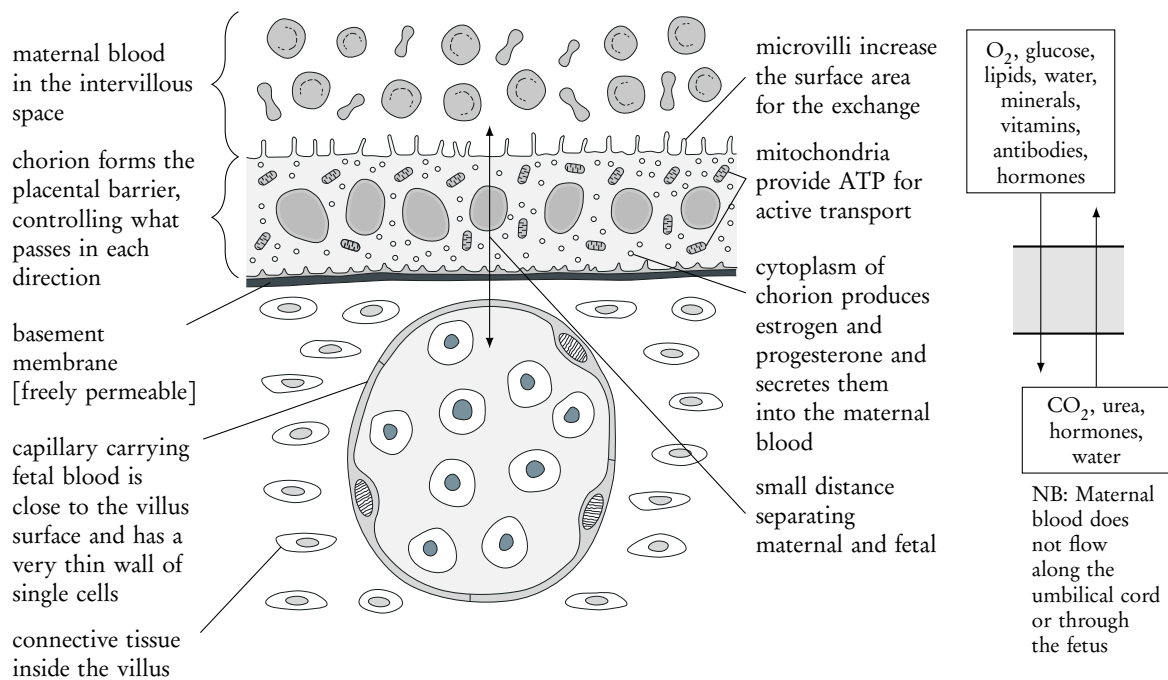


Figure 10.14: Exchange of materials across the placenta.

