### TAYLOR'S UNI. COLLEGE CHEMISTRY (9701) A Level

### APPLICATION CHEMISTRY: BIOCHEMISTRY

(Part 4 – ATP & Metals in Biological Systems)

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- The nucleotide, adenosine triphosphate (ATP), has a crucial role to play in making energy available for
- metabolic reactions in all living organisms.
- Short term energy source for cellular activity.
- In animal cells this nucleotide is synthesised in the mitochondria of the cell.

- The molecule consists of three phosphate groups linked in a short chain and covalently bonded to the hydroxyl group of a sugar, ribose.
- The last part of the molecule is adenine, an organic base.
- The breakdown of ATP is an exothermic reaction and this released energy is used by enzymes to power the catalysed reactions.
- ATP is hydrolysed to ADP (adenosine diphosphate) and an inorganic phosphate ion (P<sub>i</sub>)

- There is a net gain of energy when the products are formed.
- The release of the end phosphate group is favoured by the repulsion between the negatively charged O atoms on the adjacent phosphate groups

- **Synthesis** of **ATP** energetically **endothermic** and this is where the energy available from the oxidation of food is needed (eg. respiration).
- Plants can make ATP from ADP and phosphate ions using energy from sunlight (photosynthesis)

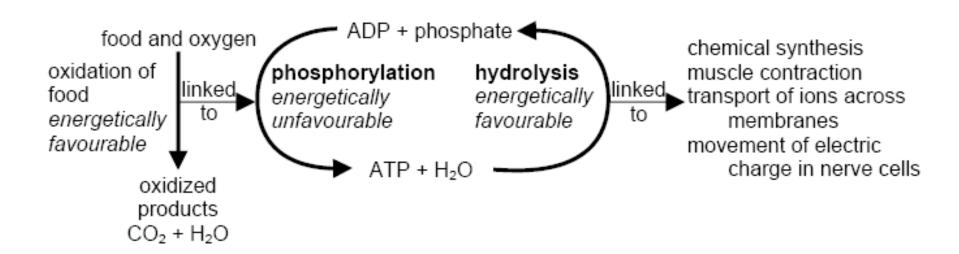


Figure 1.46 – the role of ATP in metabolism

- If cellular metabolic activity is to be controlled then spontaneous reactions (hydrolysis of ATP) must not occur in the absence of enzymes.
- Although the hydrolysis of ATP to ADP is energetically favourable, the activation energy for the reaction is high.
- This means that spontaneous hydrolysis of ATP without the presence of an enzyme does not occur.

### Metals in Biological Systems

#### (1) Iron and the haem proteins

- The Fe<sup>2+</sup> ions that bind oxygen to the haemoglobin.
- Each haem group can bind one oxygen molecule, and each of the four haem groups binds oxygen simultaneously, so the overall reaction is:

- Fe<sup>2+</sup> ions act as the centres of complex ions; the **ligands** being the **haem group**, the protein chain, and the attached oxygen molecule
- The **binding** of the oxygen is **reversible** so that it can be 'delivered' to the tissues of the body where it is required.

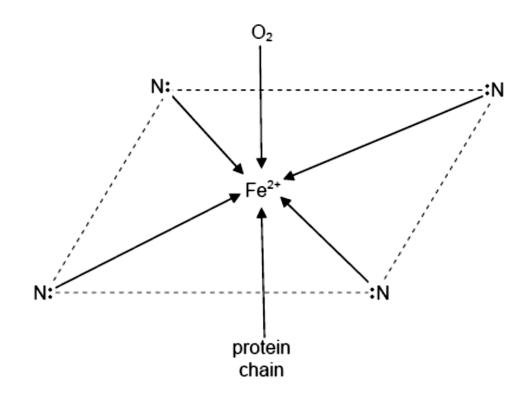


Figure 1.47 – the complex ion in haemoglobin

- Oxygen ligand can be replaced by another ligand that binds more strongly.
- In carbon monoxide poisoning the CO molecules replace the oxygen in oxyhaemoglobin.
- Haem is also involved in the functioning of other proteins such as the cytochromes present in the mitochondria.
- These proteins are responsible for the production of ATP in the final stage of the respiration process.
- Redox processes involving electron transfer are crucial at this stage and the ability of the Fe<sup>2+</sup> ions to form Fe<sup>3+</sup> ions relatively easily, and vice versa, is important here

#### (2) Zinc as an enzyme cofactor

- Carbonic anhydrase, present in our red blood cells, it is responsible for the removal of carbon dioxide from the blood, producing hydrogen carbonate ions.
- Key to the activity of this enzyme is the zinc ion (Zn²+)
  present in the active site of the enzyme.
- The zinc is bound to the enzyme as part of a complex using nitrogen atoms on the protein chain.
- Water is also bound to the zinc ion. Since the zinc ion has a high charge density it assists the breakdown of this water molecule into an H<sup>+</sup> and an OH<sup>-</sup> ion.

- The hydroxide ion act as an nucleophile and attack the carbon dioxide molecule 

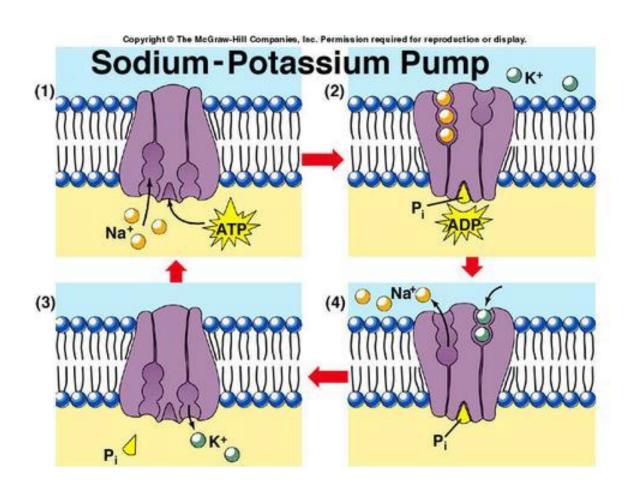
  the hydrogen carbonate ion is released from the active site.
- → a further water molecule binds to the zinc and the catalytic cycle begins again.

$$CO_2$$
 +  $OH^- \rightarrow HCO_3^-$ 

### Sodium and potassium ion transfer across cell membranes

- Within cells: [Na+] is lower, [K+] is higher, than the surrounding liquid outside.
- When a nerve is stimulated sodium ions pour into the nerve cell.
- When this 'signal' has passed the Na<sup>+</sup> and K<sup>+</sup> ion concentrations have to be restored to normal.
- The sodium being transported out of the cell once again.
- The energy needed for this active transport come from the hydrolysis of ATP (enzyme for this reaction Na<sup>+</sup>, K<sup>+</sup> -ATPase or 'sodium-potassium pump')

 These enzyme molecules sit across the membrane with parts of the protein exposed on the outer and inner surfaces.



- (1) three Na<sup>+</sup> ions and an ATP molecule bind to the inner protein surface of the enzyme.
- (2) ATP is hydrolysed, with the Pi binding to the protein.
- Enzyme changes shape so that the Na<sup>+</sup> ions move to the outside surface
- (3) Two K<sup>+</sup> ions attach after Na<sup>+</sup> released.
- (4) Two K<sup>+</sup> ions released into the cell.
- When another ATP binds, process is repeated.

#### The potassium specific channel

- There are also specific water and ion channels in cell membranes.
- their selectivity is dependent on the hydration and size of the ions concerned.
- the aqueous K<sup>+</sup> ions (K<sup>+</sup>(aq)) must lose their hydration shells before they can pass through the channel → linking instead to oxygen atoms in certain R groups of the protein.
- The selectivity of the channel depends on the distances between the oxygen atoms in the protein side-chains and the K+ ions.
- The smaller Na<sup>+</sup> ions will not 'fit' the channels as the distances are too great for the complex to form.

# The potassium specific channel

- The enthalpy required to lose the hydration shell around the ions is compensated for by that given out when the new association is formed with the protein.
- The K<sup>+</sup> ions pass through the channel and then reassociate with water on the other side.
- The selectivity of the channel depends on the distances between the oxygen atoms in the protein side-chains and the K<sup>+</sup> ions.
- The smaller Na<sup>+</sup> ions will not 'fit' the channels as the distances are too great for the complex to form.

## Toxic trace metals in the environment

- Lead and mercury interfere with enzyme function by disrupting the disulphide bridges involved in protein tertiary structure.
- The presence of certain salts disrupt Van der Waals' forces between non-polar side-chains that gives rise to the protein tertiary structure.
- lons such as Li<sup>+</sup>, Mg<sup>2+</sup>, and Ca<sup>2+</sup>, together with heavy metal ions such as lead or mercury, have all been shown to interfere with Van der Waals' interactions.

### Later Symptoms of Lead Poisoning Memory Problems Nausea Kidney Problems Weight Loss Constipation Weak Wrists or Ankles

## Toxic trace metals in the environment

- Mercury contamination is perhaps the most notorious case of heavy metal poisoning.
- Mercury can enter the food chain by a number of routes:
  - in waste water discharged into rivers from factories that use mercury compounds in their processes,
  - mercury compounds have been used as fungicides and these can be washed off crops into the soil,
  - mercury compounds have been used to treat timber and again they can be washed into rivers and streams,

## Toxic trace metals in the environment

 Mercury cathode cell is one which is used in the large scale production of sodium hydroxide – again any leakage of mercury is dangerous as micro-organisms can convert mercury salts into organomercury compounds e.g. methylmercury salts, and these can be ingested by water-borne organisms.

• Effects of mercury toxicity: loss of muscle coordination and mental function.