# **Cellular Respiration & Glycolysis**

## **Cellular Respiration**

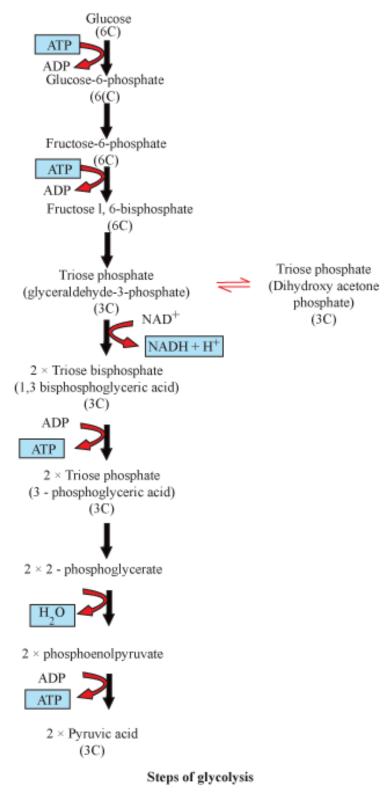
- Mechanism of breakdown of food materials within the cell to release energy, and the trapping of this energy for ATP synthesis.
- Respiratory substrates: Compounds oxidised during respiration; usually carbohydrates; these can also be proteins, fats or organic acids
- ATP: Energy currency of a cell; broken down whenever energy is needed
- Plants do not need specialised organs for respiration because:
- Gaseous exchange occurs in each part of a plant.
- Gaseous exchange is not a much-needed factor. During photosynthesis, the need for gaseous exchange is met.
- Each living cell is located close to the surface of a plant. So the distances that the gases must diffuse are not great.
- Equation for respiration:

 $C_6H_{12}O_6 + 6O_2 \rightarrow 6CO_2 + 6H_2O + Energy$ 

# Glycolysis (Glycos - Sugar, Lysis - Splitting)

- Breakdown of glucose to pyruvic acid by partial oxidation
- Scheme given by Embden, Meyerhof and Parnas
- Common pathway for aerobic and anaerobic cellular respiration
- Occurs in the cytoplasm of a cell
- Present in all living organisms
- In plants, sucrose is converted into glucose.
- Sucrose  $\longrightarrow$  Glucose  $\rightarrow$  Enters the glycolysis

- A chain of 10 reactions converts glucose into pyruvate.
- Hexokinase: Enzyme that phosphorylates glucose to produce glucose 6 phosphate
- 2 ATPs are utilised in two steps:
- Glucose  $\rightarrow$  Glucose -6 phosphate (1 ATP)
- Fructose 6 phosphate  $\rightarrow$  fructose 1, 6 bisphosphate (1 ATP)
- Fructose 1, 6 bisphosphate splits into glyceraldehyde 3 phosphate and dihydroxy acetone phosphate.
- Glyceraldehyde 3 phosphate converts into two molecules of 1, 3 bisphosphoglycerate (BPGA), with subsequent conversion of NAD<sup>+</sup> to NADH + H<sup>+</sup>.
- 4 ATPs are yielded in two steps:
- BPGA  $\rightarrow$  PGA (1 × 2 = 2 ATPs)
- Phosphoenol pyruvate  $\rightarrow$  Pyruvic acid (1 x 2 = 2 ATPs)
- ATPs produced directly = 4 (produced) 2 (consumed) = 2 ATPs
- Net ATPs Produced = 2 (NADH+ H<sup>+</sup>) = 6 ATPs + 2 (Directly synthesised) = 8 ATPs



- The pyruvate, so produced, may undergo:
- Lactic acid fermentation

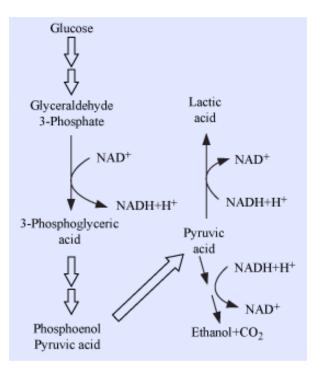
- Alcoholic fermentation
- Aerobic respiration (Krebs cycle)

# Fermentation

- Fermentation: Incomplete oxidation of glucose under anaerobic conditions
- In yeast fermentation:
- Pyruvic acid → Ethanol + CO<sub>2</sub>
- Enzymes involved Pyruvic acid decarboxylase, Alcohol dehydrogenase
- In bacterial fermentation:
- Pyruvic acid  $\rightarrow$  Lactic acid
- Enzyme involved Lactate dehydrogenase
- Similar reaction occurs in animal muscles in anaerobic conditions, say during exercise.
- Reducing agent in both reactions is NADH.

 $\mathsf{NADH} + \mathsf{H^+} \to \mathsf{NAD^+}$ 

- Only 7% of energy of glucose is released during fermentation.
- Process can be hazardous as alcohol or acid is produced. Yeasts poison themselves to death when alcohol concentration reaches about 13%.



# Tricarboxylic Acid Cycle

## **Aerobic Respiration**

- Site: Mitochondria
- Events:
- **TCA cycle** (in the mitochondrial matrix) complete oxidation of pyruvate by stepwise removal of all hydrogen atoms, which leaves three molecules of CO<sub>2</sub>
- Electron Transport Chain and Oxidative phosphorylation (in the inner membrane of the mitochondria) electrons removed as a part of hydrogen atoms are passed on to molecular oxygen, with the simultaneous synthesis of ATP

## Formation of Acetyl Coenzyme A

• The product of glycolysis, i.e., pyruvate, on entering the mitochondrial matrix, undergoes *Oxidative Decarboxylation*, thereby producing acetyl CoA which enters Krebs cycle.

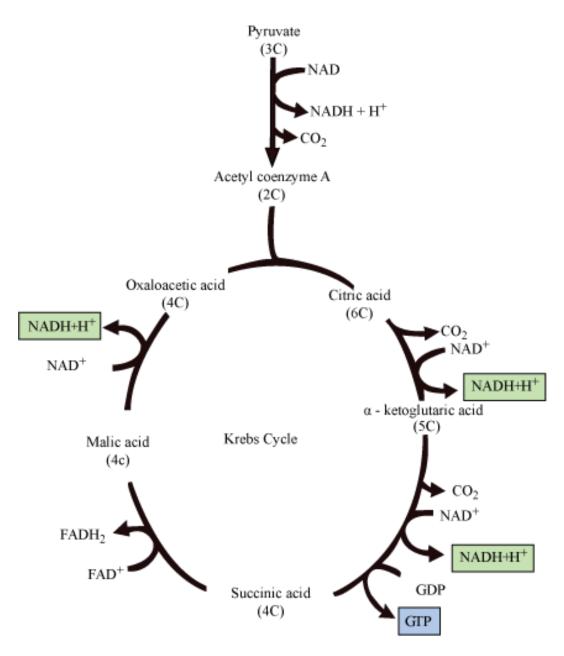
Pyruvic acid + CoA + NAD<sup>+</sup>  $\xrightarrow{Mg^{2+}}$  Acetyl CoA + CO<sub>2</sub> + NADH + H<sup>+</sup>

Krebs cycle (Tricarboxylic acid cycle; TCA)

- Acetyl group condenses with oxaloacetic acid and water to yield citric acid (catalysed by the enzyme citrate synthase)
- Citrate isomerises into isocitrate
- Two successive decarboxylation steps follow, leading to the formation of α ketoglutaric acid, followed by succinyl CoA
- This is followed by the conversion of succinyl CoA into succinic acid. During this process, GDP is converted into GTP (substrate level phosphorylation).
- In a coupled reaction, GTP is converted into GDP, simultaneously synthesising ATP from ADP.
- Conversion of one molecule of pyruvate into acetyl CoA yields 1 molecule of CO<sub>2</sub> and 1 NADH.
- One Kreb's cycle yields 2 CO<sub>2</sub> + 3 NADH + 1 FADH<sub>2</sub> + 1 ATP
- Overall equation:

Pyruvic acid + 4NAD<sup>+</sup> + FAD<sup>+</sup> + 2H<sub>2</sub>O + ADP + Pi  $\xrightarrow{\text{Mitochondrial}}$  3CO<sub>2</sub> + 4NADH + 4H<sup>+</sup> + FADH<sub>2</sub> + ATP

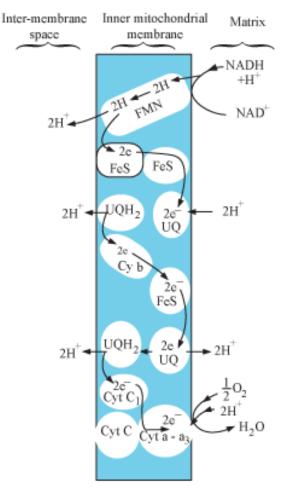
- For continued oxidation of acetyl CoA in TCA 2 things are required:
- Continued replenishment of oxaloacetic acid
- Regeneration of NAD<sup>+</sup> and FAD<sup>+</sup> from NADH and FADH<sub>2</sub> respectively.



## **Electron Transport Chain and Oxidative Phosphorylation**

## **Electron Transport Chain (ETS)**

- NADH and FADH<sub>2</sub> are oxidised to release the energy stored in them.
- Electrons are passed from one carrier to another, and finally to oxygen, resulting in the formation of water.



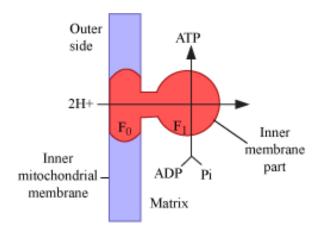
Electron Transport System (ETS)

- Electrons produced by NADH in the mitochondrial matrix are oxidised by Complex I (NADH dehydrogenase).
- Electrons are then transferred to ubiquinone, located in the inner mitochondrial membrane. Ubiquinone receives reducing equivalents through **Complex II** (FADH<sub>2</sub>).
- Reduced ubiquinone (ubiquinol) is then oxidised by the transfer of electrons from **Complex III** (cytochrome *bc*<sub>1</sub> complex) to cytochrome *c*.
- Cytochrome *c* transfers the electrons between Complex III and Complex
  IV (Cytochrome *c* oxidase complex consists of cyt *a* and *a*<sub>3</sub>, along with two copper centres).
- In the course of passing from one carrier to another, electrons couple with Complex V (ATP synthase) and produce ATP.
- Oxidation of 1 NADH produces 3 ATPs. Oxidation of 1 FADH<sub>2</sub> produces 2 ATPs.

• Role of oxygen in the terminal stage of ETS: It acts as the final hydrogen acceptor; removes hydrogen from the process and drives the whole process

#### **Oxidative Phosphorylation**

- Production of proton gradient needed for the production of ATP is provided by the energy of oxidation-reduction reaction. Therefore, the process is called oxidative phosphorylation.
- Complex V (ATP synthase) is involved. It has two major components
- F<sub>0</sub> integral membrane protein; forms a channel through which H<sup>+</sup> cross the inner membrane
- F<sub>1</sub> passage of H<sup>+</sup> induces conformational changes in F<sub>1</sub>, which forms a site for synthesis of ATP from ADP; for each ATP produced, 2H<sup>+</sup> pass through F<sub>0</sub>, down the electrochemical proton gradient



# **Respiratory Balance Sheet and Respiratory Quotient**

## **Respiratory Balance Sheet**

- It gives the net gain of ATP for every 1 molecule of glucose oxidised.
- Certain assumptions are made for calculating the net gain of ATP:
- An orderly pathway is followed Glycolysis, followed by TCA cycle, followed by ETS
- NADH synthesised during glycolysis enters the mitochondria to undergo oxidative phosphorylation.
- Except glucose, no other substrate enters the pathway at any stage.

- Intermediates do not synthesise any other compound in the pathway.
- There can be a net gain of 36 ATPs during aerobic respiration of 1 molecule of glucose. Glucose +  $6O_2$  + 36ADP +  $36Pi \rightarrow 6CO_2$  +  $42H_2O$  + 36ATP

Fermentation	Aerobic Respiration
1. Partial breakdown of glucose	1. Complete breakdown of glucose into $CO_2$ and $H_2O$
2. Net gain of only 2 molecules of ATP	2. Net gain of 36 molecules of ATP
3. Here, oxidation of NADH to NAD⁺ is a slow reaction	3. Here, oxidation of NADH to NAD <sup>+</sup> is a vigorous reaction

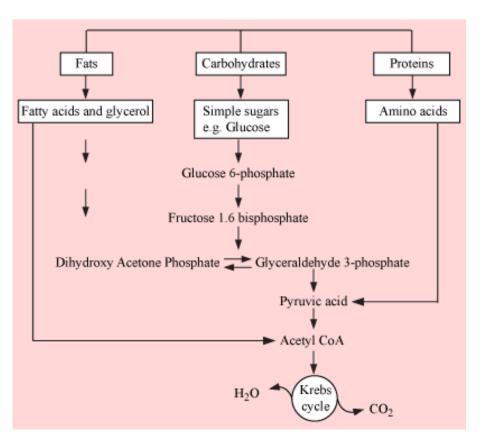
#### Comparison between fermentation and aerobic respiration

#### **Amphibolic Pathway**

- Favoured substrate for respiration is glucose. All carbohydrates first convert into glucose to enter the pathways.
- Other substrates do enter the respiratory pathways, but not during the first stage.
- Fats:
- Fats  $\rightarrow$  Glycerol + Fatty acid
- Fatty acids → Acetyl CoA Acetyl CoA enters the pathway
- Glycerol  $\rightarrow$  PGAL
- Proteins:

• Proteins deamination Amino acids

• Enter intermediate stages of TCA



- Amphibolic pathway: Involved in both anabolism and catabolism
- Fatty acids break into acetyl CoA to enter the respiratory pathway (Anabolism).
- Acetyl CoA is removed from the respiratory pathway whenever fatty acids need to be synthesised (catabolism).
- Thus, respiratory intermediates form a link during anabolism and catabolism.

#### **Respiratory Quotient (RQ)**

• Ratio of the volume of CO<sub>2</sub> evolved to the volume of O<sub>2</sub> consumed during respiration is called the respiratory quotient (RQ).

Volume of CO<sub>2</sub> evolved

 $RQ = Volume of O_2 consumed$ 

- Depends upon the type of respiratory substrate
- RQ = 1 (When carbohydrate is used as substrate)

•  $C_6H_{12}O_6 + 6O_2 \rightarrow 6CO_2 + 6H_2O + Energy$ 

$$RQ = \frac{\frac{6CO_2}{6O_2}}{1.0} = 1.0$$

• RQ < 1 (When fat is used as respiratory substrate)

E.g. When fatty acid, tripalmitin is used, RQ = 0.7

- RQ  $\approx$  0.9 (When protein is used as substrate)
- RQ is infinity in anaerobic respiration as CO<sub>2</sub> is evolved, but O<sub>2</sub> is not utilised.