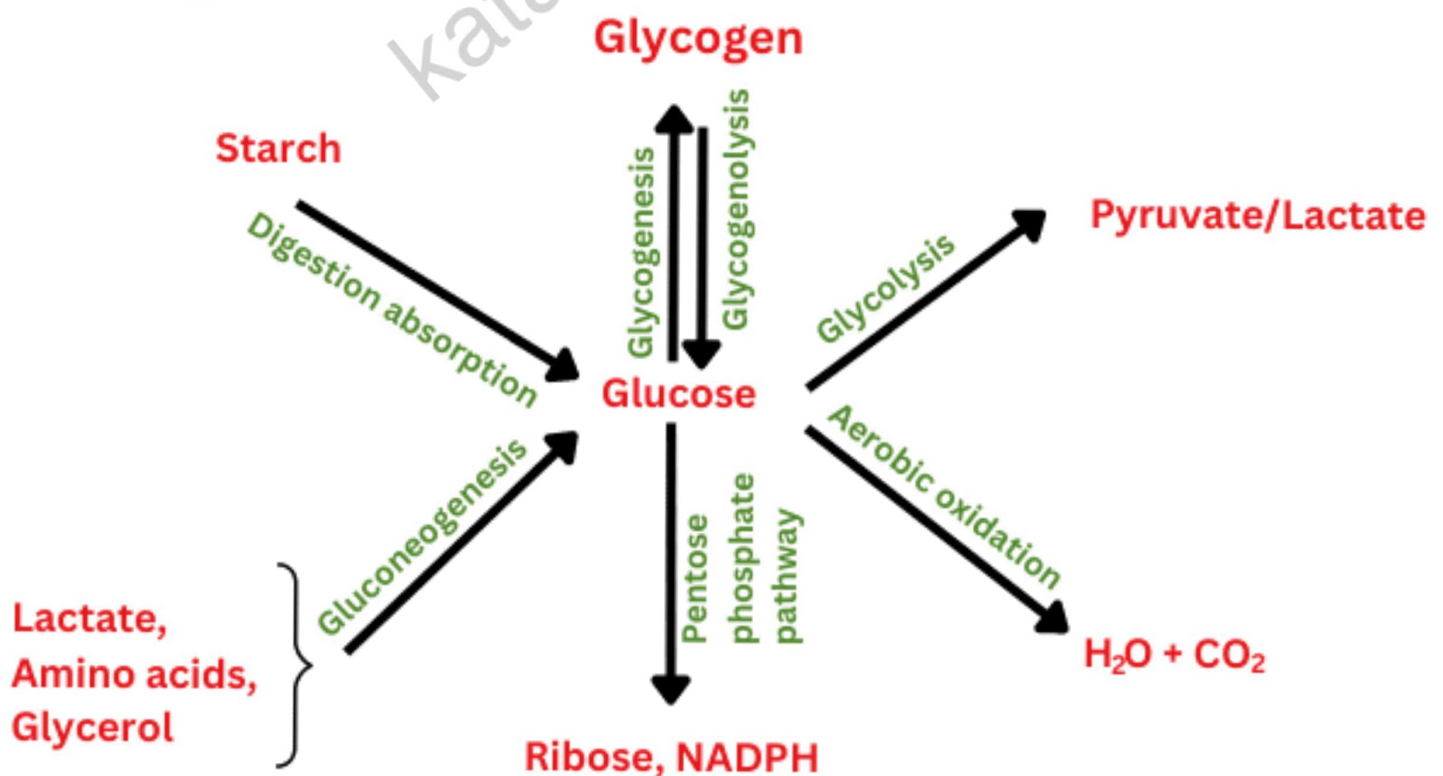


★ Carbohydrate Metabolism :-

- Biochemical processes involves in Synthesis, breakdown and inter-conversion of Carbohydrates in living organism are Collectively Called as Carbohydrate Metabolism.
- Glucose is the essential molecule of Carbohydrate Metabolism which participates in various metabolic pathways.
- Insulin is the primary metabolic hormone Synthesised in pancreas and regulates glucose level in blood.

Carbohydrate metabolism

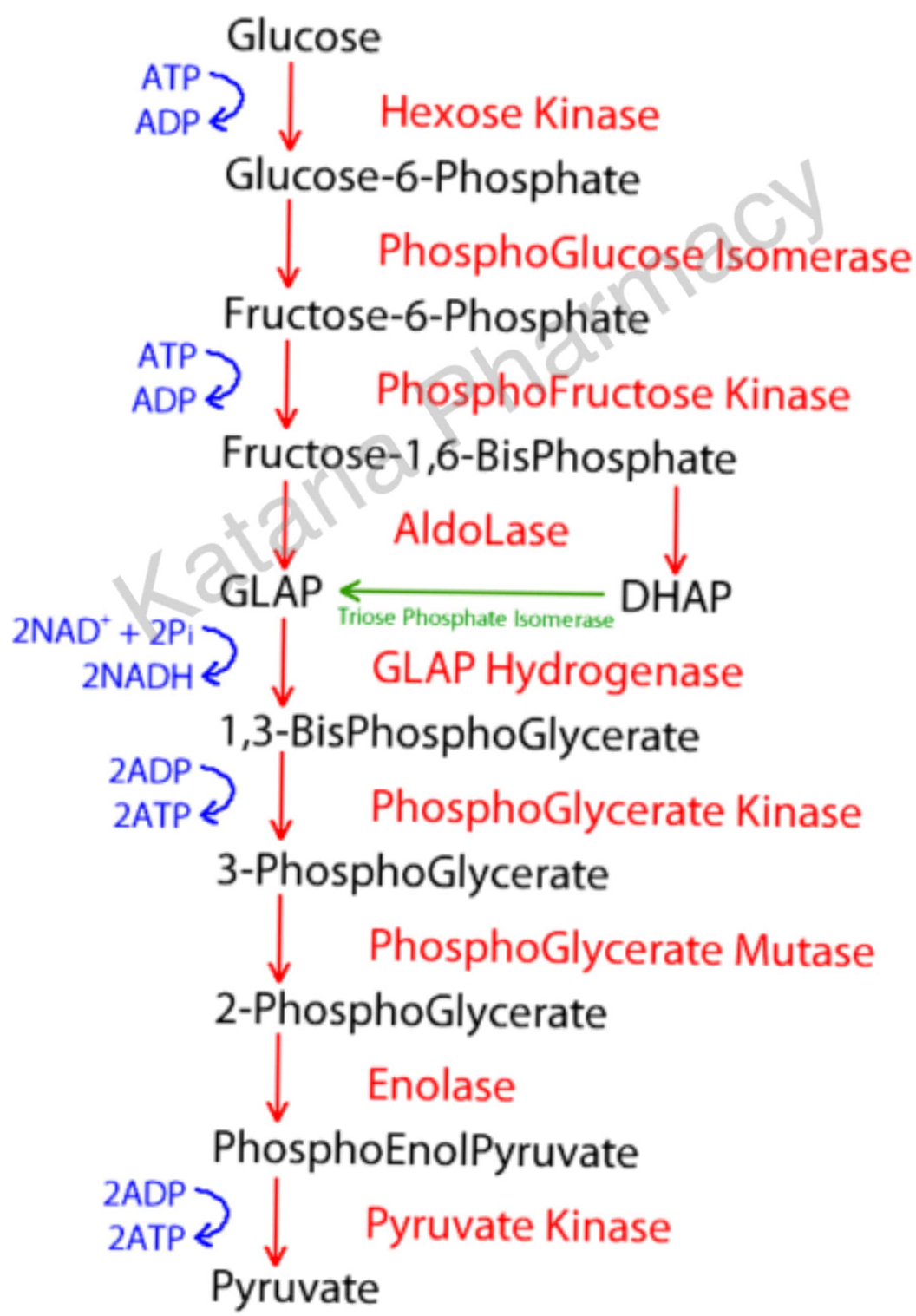


Major Pathways of Carbohydrate Metabolism :-

① Glycolysis :-

Glycolysis is an important pathways of Carbohydrate metabolism and occurs Cytosol of a living cell.

- In glycolysis one mol glucose yield two moles of Pyruvate, ATP and NADH.



Energetics :-

- Net gain in glycolysis is 2 ATP, 2 pyruvate and 2 NADH_2 molecule.
- Glycolysis is a low energy process as pyruvate, and NADH_2 molecule captures much of the energy.

Table: Input and output of material during Glycolysis

<u>Total input</u>	<u>Total output</u>
1 Molecule of glucose (6c)	2 molecule of pyruvic acid (2x3c)
2 ATP	4 ATP
4 ADP	2 ADP
2 x NAD	2 x NADH_2
2 x P_i	2 x H_2O

- Total 8 ATP generate during one cycle of glycolysis.

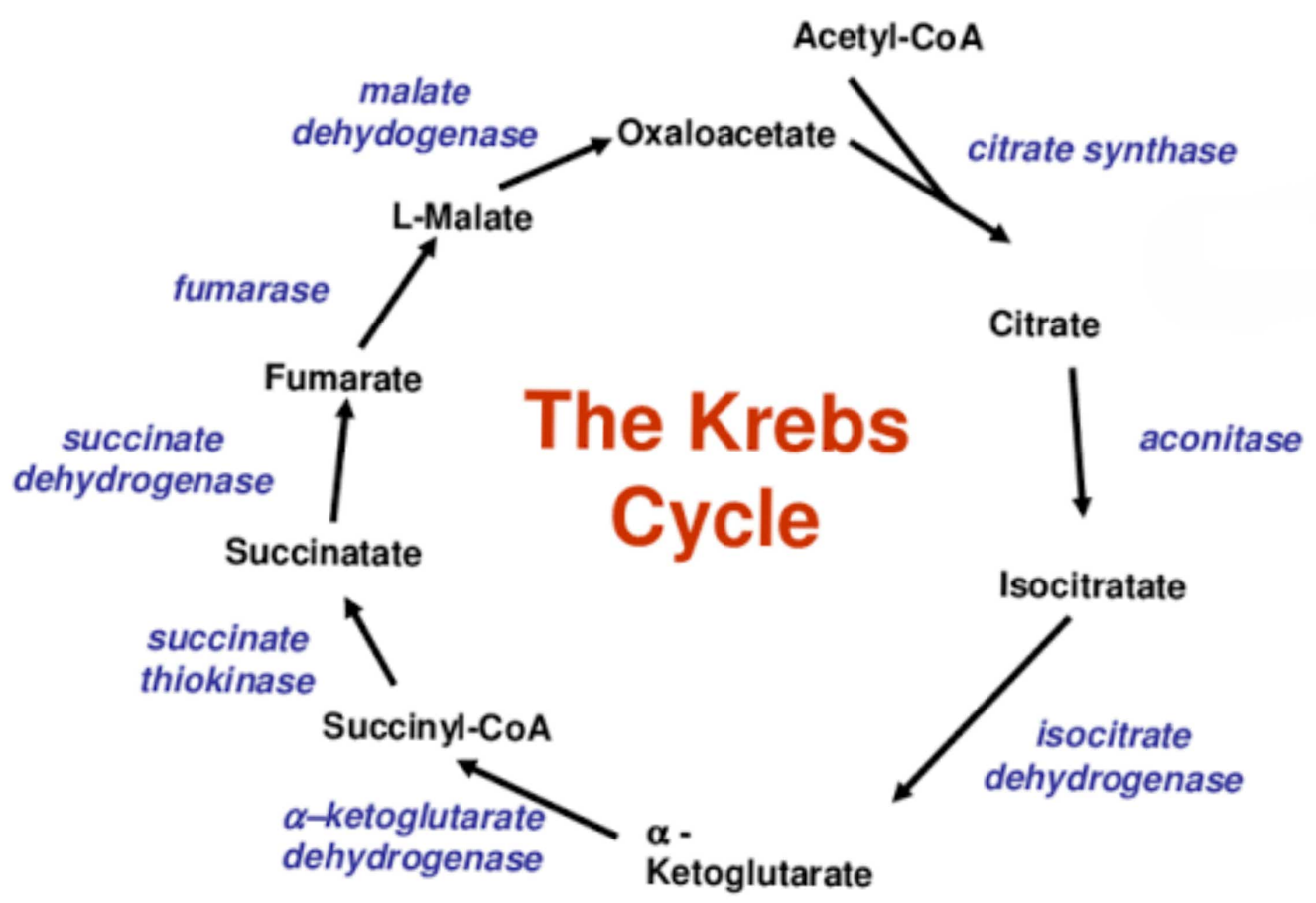
Significance :-

- ① In glycolysis pathway, glucose is oxidised to yield pyruvate
- ② Enzyme involve in glycolysis pathway are present in Cytosol of a cell.
- ③ All cell utilise glucose for energy
- ④ In a single pathway of glycolysis two molecule of ATP and NADH are generated.
- ⑤ Energy produce by anaerobic glycolysis provides reduce oxygen supply in cell.
- ⑥ Regulation of glycolysis is important to maintain ATP balance.

② Citric acid Cycle:-

- The citric acid Cycle (Krebs Cycle or tricarboxylic acid - TCA Cycle) is the most important metabolic pathway for the energy supply to the body.
- ATP is synthesized in Krebs Cycle. Citric acid Cycle essentially involves the oxidation of acetyl CoA to CO_2 and H_2O .

Pathway



- Once pyruvate undergoes oxidative decarboxylation to form acetyl CoA, the acetyl CoA then enters the Citric acid Cycle, also known as the Krebs Cycle.
- The Citric acid Cycle is a series of eight reactions that break down acetyl CoA, the product of pyruvate decarboxylation, to yield high-energy electron-carrier molecules such as NADH and $FADH_2$.
- The purpose of the Citric acid Cycle is to harvest the "high energy" electron from the carbon fuel source that ultimately comes from glucose.

Step 1: Oxaloacetate and acetyl CoA react to form citryl CoA, which then reacts with water to form citrate. The purpose of this step is to create a molecule that will ultimately undergo decarboxylation.

Step 2: The hydroxyl group on citrate is not positioned correctly to undergo a decarboxylation, therefore citrate is transformed into isocitrate by an enzyme called aconitase.

Step 3: This is the first oxidation-reduction reaction in the Citric acid Cycle, hence we first reduce NAD^+ into NADH and then produce the α -ketoglutarate via an oxidative decarboxylation reaction.

Step 4: - This is the second oxidative decarboxylation reaction where we form a four-carbon molecule, releasing CO_2 and forming NADH.

Step 5:- In this step, we transfer a phosphate group from Succinyl CoA into a GDP. to form GTP. This is the only step in the citric acid cycle that produces with a high phosphoryl transfer potential.

Step 6-8:- The final three step of 1 citric cycle involve the regeneration of the oxaloacetate molecule.

- In this process of the regeneration of oxaloacetate, also form FADH_2 (step 6), NADH (step 8) and use water (step 7)

Product = 4CO_2 , 6NADH , 2FADH_2 , 2GTP , 4H^+ , 2CoA .

③ Hmp Shunt :-

- Hmp (Hexose mono phosphate) Shunt is also known as pentose phosphate pathway.
- It is an alternative pathway for glucose oxidation and is a multi-cycle process.
- Enzyme of Hmp shunt are located in Cytosol.

pathway :-

Reaction of the pathway -

Reaction of Hmp pathway is divided into Two phases .

1. oxidative phase
2. Non-oxidative phase

1. oxidative phase :-

Step-1 :- Glucose-6-phosphate Convert into 6-phosphogluconolactone in presence of enzyme Glucose-6-phosphate dehydrogenase . (Rate limiting step)

Step 2 :- 6-phosphogluconolactone Convert into 6-phosphogluconate in presence of enzyme Gluconolactone hydrolase.

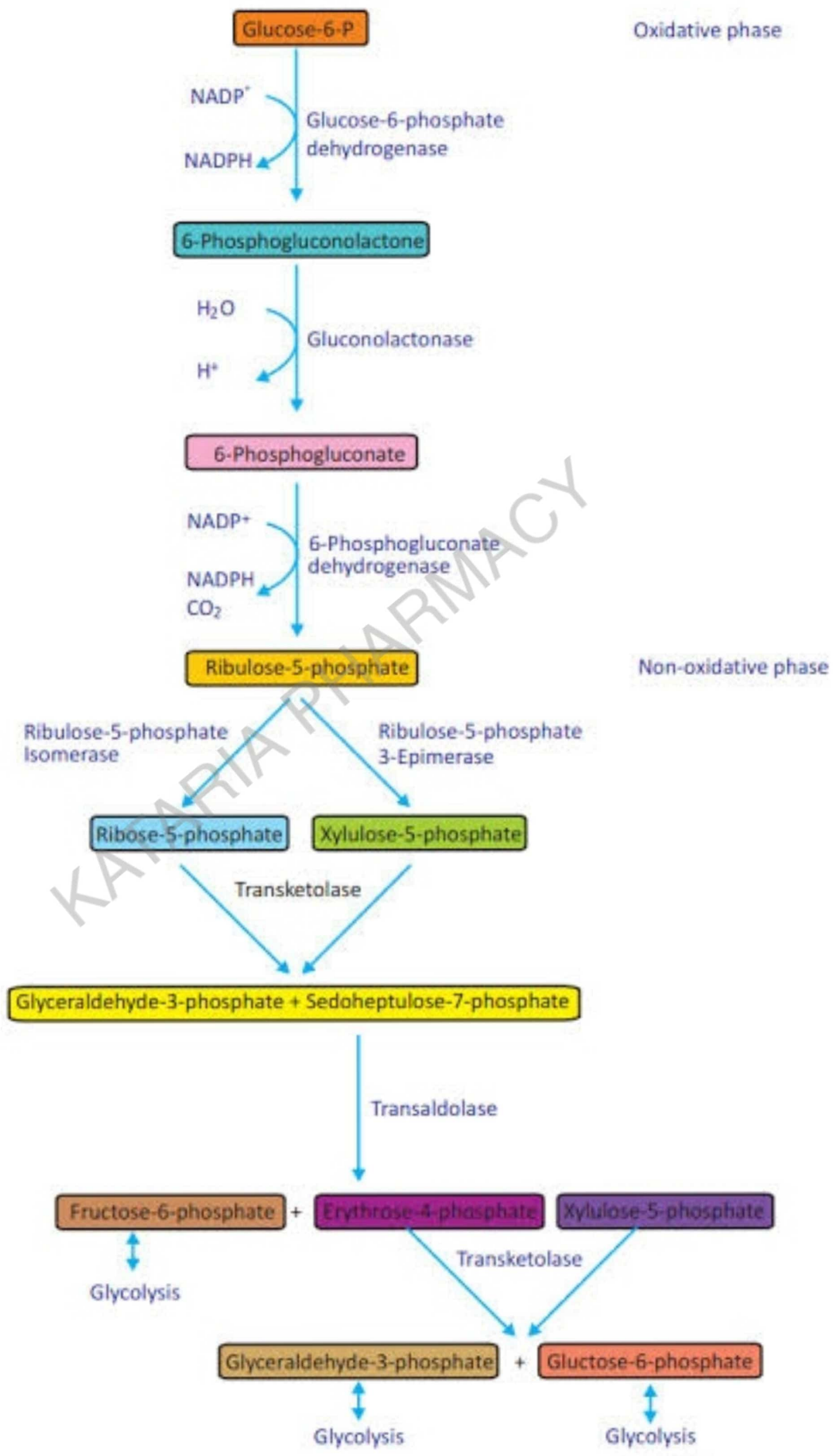
Step 3 :- NADPH is again generated.

6-phosphogluconate Convert into Ribulose-5-phosphate in presence of enzyme 6-phosphogluconate dehydrogenase and 6-molecule of CO_2 is released.

② Non-oxidative phase:-

Step 4:- Termination -

Ribulose-5-phosphate Convert in *epimerase* → xylulose-5-phosphate
Isomerase → Ribulose-5-phosphate



Step 5 :- Transketolase Reaction

(27)



Step 6 :- Transaldolase Reaction -

Sedoheptulose-7-phosphate and Glyceraldehyde-3-phosphate Convert into Erythrose-4-phosphate and Fructose-6-phosphate in presence of enzyme Transaldolase.

Significance :-

- Free radical Seavenging
 - The free radicals (super oxide, hydrogen peroxide) are Continuously produce in all Cells.
- Hmp Shunt is unique in generating two important product - Pentose and NADPH
- NADPH is required for the bio Synthesis of fatty acids and Steroids.
- NADPH is required by the RBC to keep the glutathione in the reduced state.

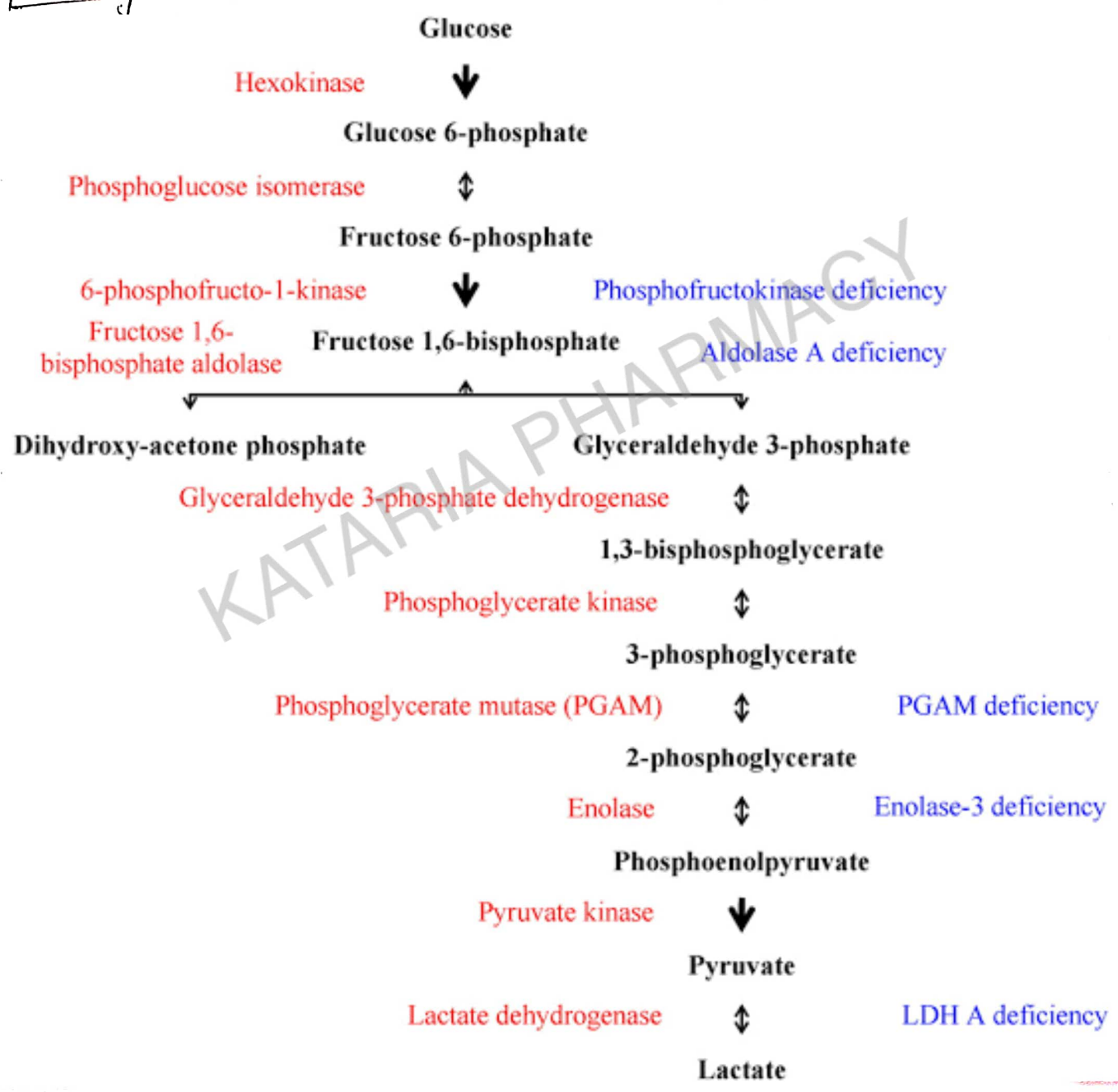
★ Glycogen Metabolism pathways:-

- In animals, glucose is stored in the form of glycogen.
- Glycogen is stored in the form of granules in cell Cytosol where most of the enzyme required for the Synthesis and breakdown are found.

Glycogenesis:-

Glycogenesis is glycogen Synthesis from glucose. This occurs in Cytosol with the help of Adenosine Triphosphate and Triphosph Uridine triphosphate.

Pathway



★ Glycogen Storage Disease (GSD) :-

- Glycogen Storage disease are genetic disorder of glycogen Metabolism occurring due to accumulation of large amount of glycogen or its metabolites into the tissue.
- Glycogen accumulates in the absence of glycogen metabolising enzyme. Not all but few of the disorder are serious.
- Glycogen Storage diseases are as follow.

① Von Gierke's Disease :-

This disease is related to the deficiency or complete absence of enzyme glucose-6-phosphate in liver, kidney and intestine.

② Pompe's Disease :-

It is a fatal disease arising due to decrease in lysosomal α -glucosidase, thus glycogen is not utilised by lysosomes and accumulates in body tissue.

③ Anderson's disease :-

In this disease, an intermediate product of glycogenesis amylopectin deposits in the liver, spleen and heart.

④ Other :-

- ① Cori's disease
- ② McArdle's Syndrome
- ③ Hers Disease

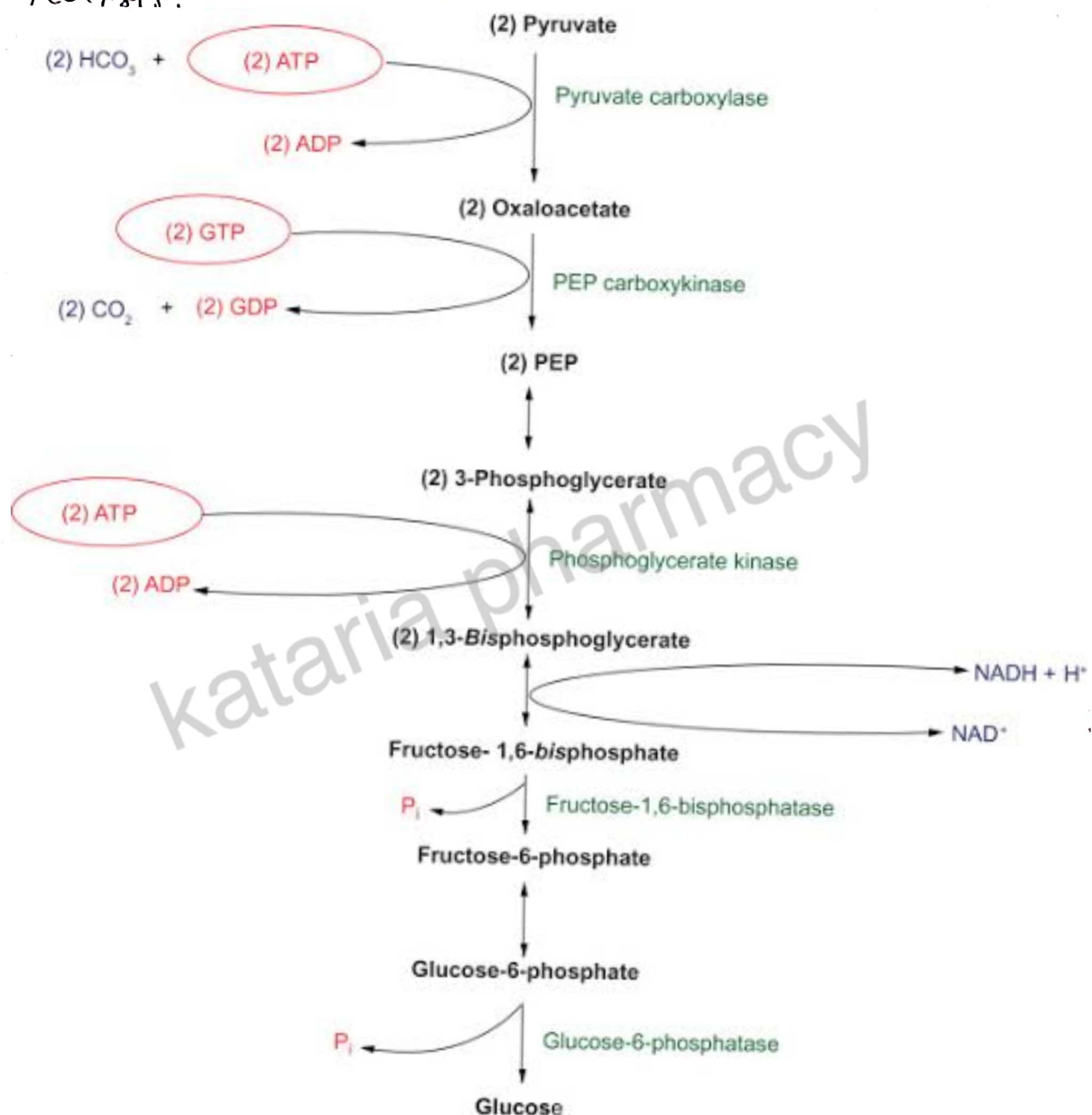
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* Gluconeogenesis:-

- Gluconeogenesis is a metabolic pathway which includes the biochemistry reaction synthesising glucose from any non-carbohydrate source such as glycerol, pyruvate, lactate, and glucogenic amino acids.
- Most of the reactions of gluconeogenesis take place in the liver and few of the reactions in the cortex part of kidney.

Pathway:-

Gluconeogenesis is almost the reversible pathway of glycolysis (pyruvate converted to glucose) but not the exact reversal of glycolysis.



Significance of gluconeogenesis :-

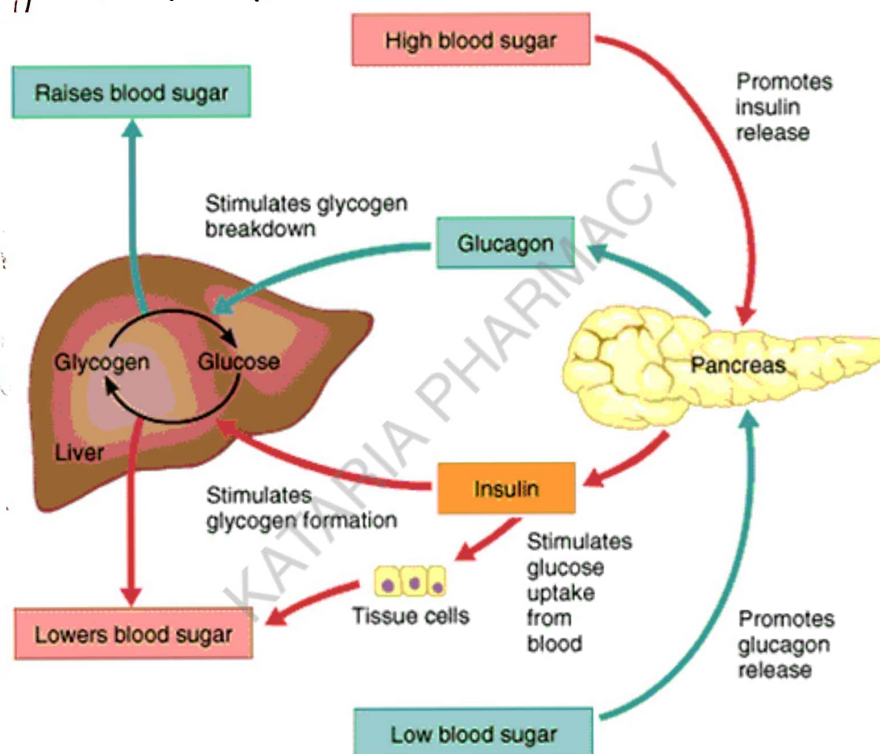
(31)

- Gluconeogenesis meets the needs of the body for glucose & maintains glucose homeostasis in absence of carbohydrate.
- Some tissue such as the brain, RBCs, lens, cornea & kidney medulla requires continuous supply of energy.
- It is used to clear the products of the metabolism of other tissue from the blood e.g. *Lactate*, *Glycerol*.
- To maintain amount of glucose.
- Removal of glycerol produced by lipolysis.

★ Hormonal Regulation of Blood Glucose Level :-

There are two categories of endocrine influences.

1. Hormone which will decrease the blood glucose levels.
Eg. Insulin
2. Hormones which will increase the blood glucose levels:
eg. Glucagon, Epinephrine, Cortisol and Glucocorticoids.

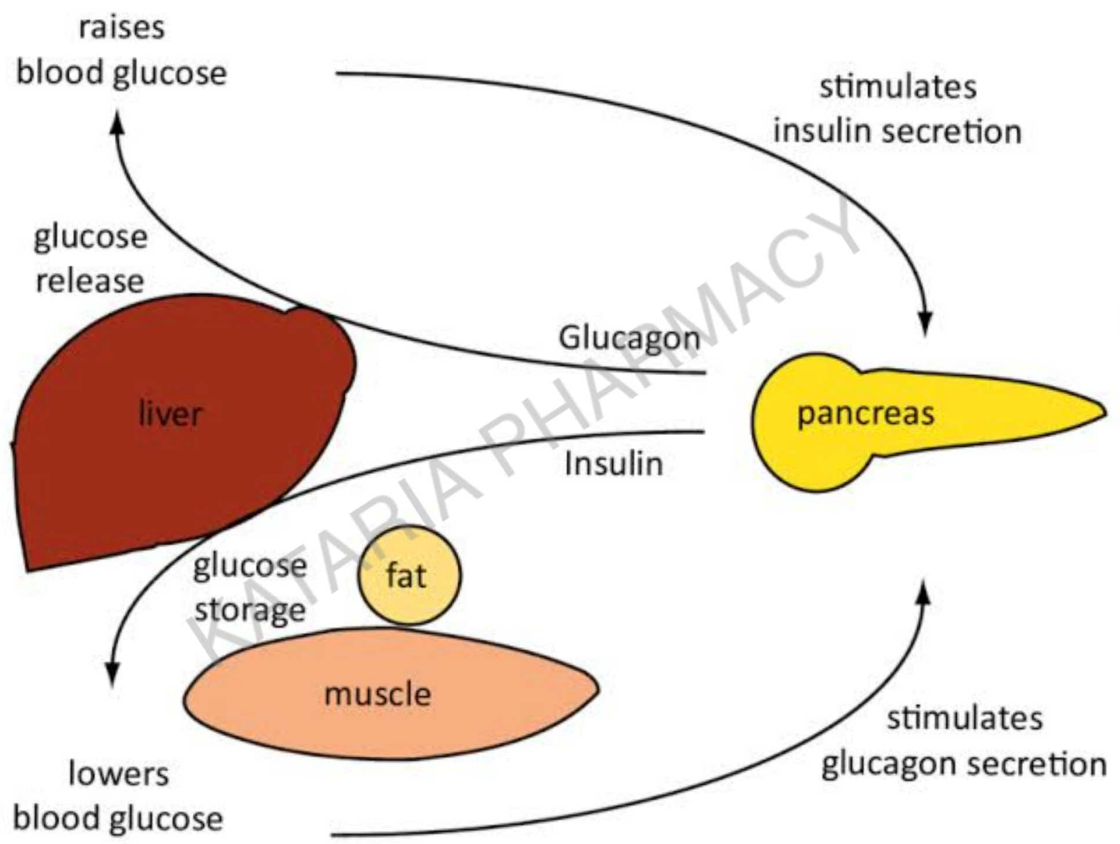


Insulin & Glucagon:

Both of these hormones are secreted by endocrine tissue of pancreas known as **Islet Cells** or **islets of Langerhans**.

- Each Islet Contains two kinds of Cells:

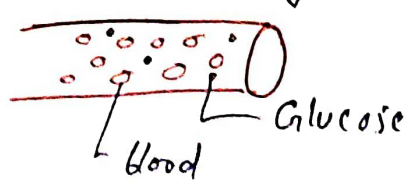
1. α -Cells (secrete glucagon) $\uparrow$ Blood glucose level.
2. β -Cells (secrete insulin) $\downarrow$ Blood glucose level



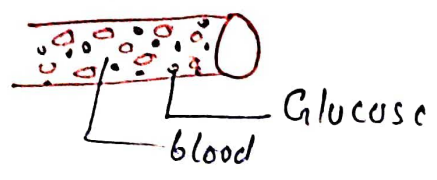
Diabetes Mellitus:

Diabetes mellitus is a metabolic disorder in which a person has high blood sugar, either because the body does not produce enough insulin.

Normal Blood glucose level



High blood glucose



Types of DM :-

① Type I DM :-

This type is characterised by destruction of β -cell (insulin producing cell) and thus insulin is required for survival.

- Effects children

- Genetic variations & auto-immune response are leading cause.

② Type II DM :-

This type is characterised by disorder of insulin action, or insulin secretion. patient have relative insulin deficiency.

- It is also known as adult onset diabetes.

Causes of DM :-

① Genetic factors $\left\{ \begin{array}{l} \rightarrow \text{Genetic defect in } \beta\text{-cell function} \\ \rightarrow \text{Genetic defects in insulin action.} \end{array} \right.$

② Environmental factor $\left\{ \begin{array}{l} \rightarrow \text{obesity associated with Modern living.} \\ \rightarrow \text{Lifestyle change (consumption of alcohol)} \end{array} \right.$

Symptoms of DM :-

① Increase thirst

② Slow healing Cuts and sores

③ Fatigue

④ Blurred vision.

⑤ Frequent urination.

⑥ Unexplained weight Loss.

★ Biological oxidation :

(24)

- In Chemical terms, removal of electrons is ~~as~~ termed as oxidation and addition of a electrons is termed as reduction.
- Thus, oxidation always goes with reduction of an electron acceptor.
- This phenomenon of oxidation-reduction also applies in biochemical system and describe biological oxidation in a living system.
- In cell during respiration, biological oxidation takes place in four stages, Carbohydrate (glucose) is used as a source of energy.

1. Glycolysis : It is the first stage in which glucose degrades into pyruvate.

2. Oxidative Decarboxylation of pyruvate :

This stage occurs for a short term in which a 2-C containing molecule is formed from the 3-C containing molecule, pyruvate with the release of CO_2 .

3. Citric acid Cycle :-

In this stage, the 2-C containing molecule is degraded into CO_2 .

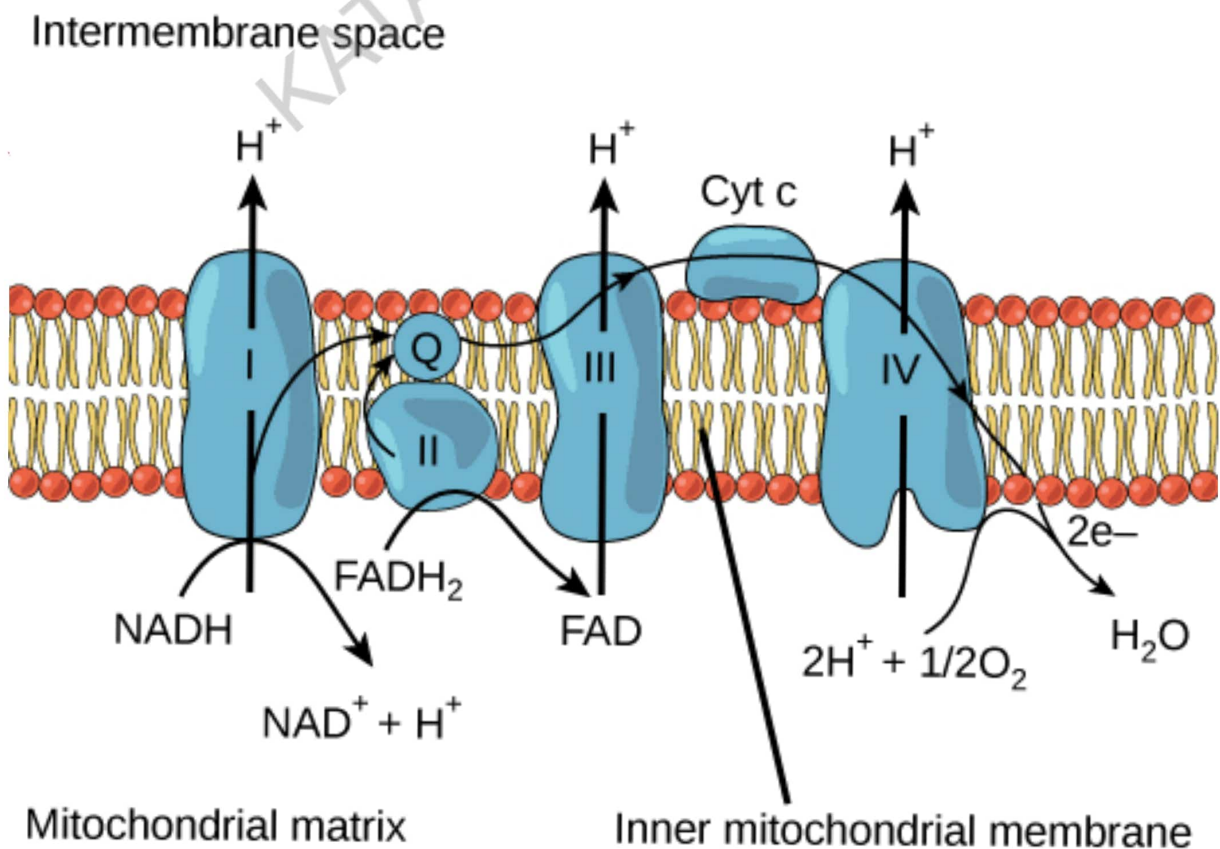
* Electron transport Chain (ETC) :-

ETC Couple a chemical reaction b/w an electron donor and electron acceptor to the transfer of H^+ ions across a membrane, through a set of mediating biochemical reaction.

- These H^+ ions are used to produce ATP.
- In ETC, the electron travel through the chain, passing from a higher to a low energy level by moving through electron-rich to electron-deficit molecule.

ETC Components :-

- 5 protein Complexes - I, II, III, IV, V
- Coenzyme Q
- Cytochrome c - a peripheral membrane protein.



Mechanism of ETC :-

(36)

The Steps involved in electron transport Chain are -

- ① The hydrogen Carriers from the citric acid Cycle enter the inner Mitochondrial Membrane folds known as Cristae.
- ② In the Cristae, the H^+ ions are oxidised by a Series of step using molecular oxygen to release energy.
- ③ A part of this energy is utilised in oxidative phosphorylation in which ATP (from ADP) and inorganic phosphate are produced.
- ④ H^+ ions or electron moves downhill i.e from a higher energy level to lower energy levels by passing from one Carrier to the next, until they reach the oxygen.
- ⑤ The oxygen accepts electrons and reduces to form water.
- ⑥ At each transfer level, Some amount of energy is released in the form of heat which is utilised for producing ATP.

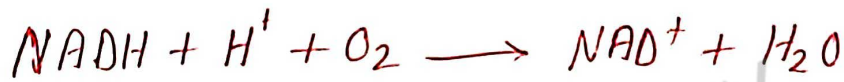
Significance :-

- ① Regenerating Electron Carrier
- ② Generating proton Gradient.

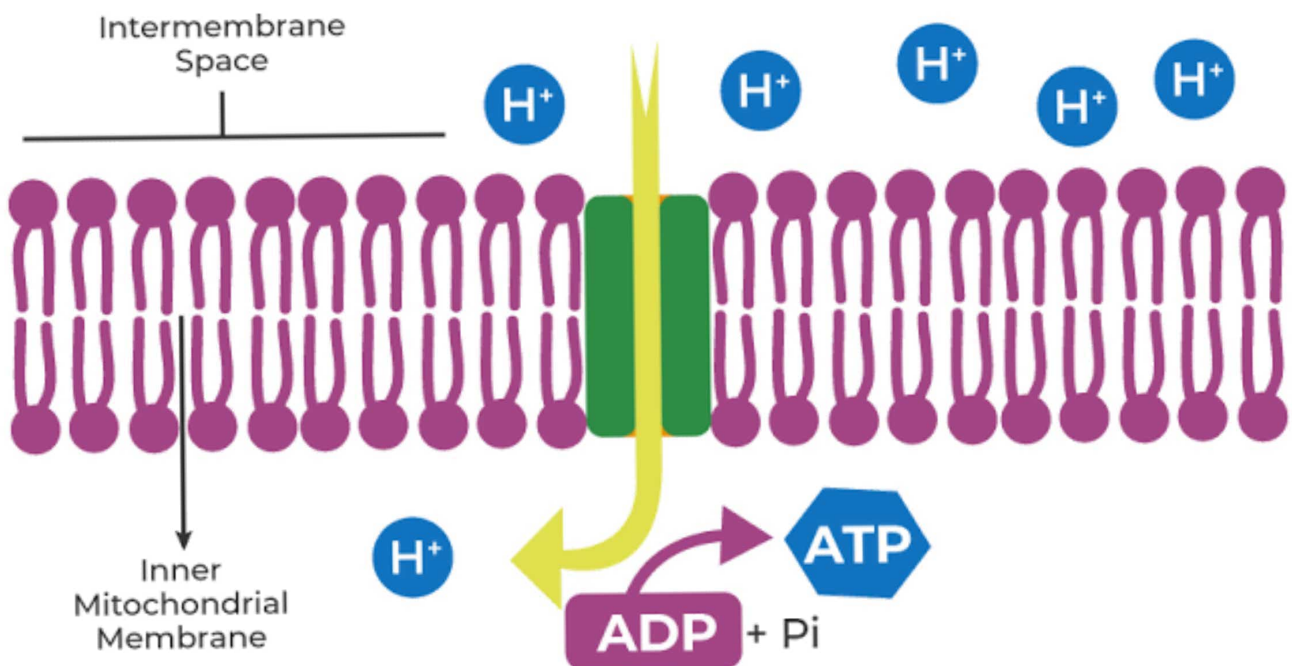
* Oxidative phosphorylation:-

- ~~oxi~~ Oxidative phosphorylation is electron transport chain or chemiosmosis.
- It is a process in which ATP is formed as a result of transfer of electrons from NADH or FADH₂ to O₂ by series of electron carriers.

A. Oxidation step : Electron transport Chain



B. Phosphorylation step:



★ Inhibitors of oxidative phosphorylation / uncouplers:-

- Electron transport within mitochondria and oxidative phosphorylation (ATP Synthesis) are combined process.
- It is also be said that oxidation and phosphorylation process proceed simultaneously.
- Some Compound have ability to uncouple these two reactions, and such compounds are known as ~~Uncop~~ uncouplers.
- The uncouplers increase permeability of the inner mitochondrial membrane towards protons (H^+) and thus inhibiting ATP formation.
- The energy generated during electron transport is dissipated in the form of heat.
- Thus, the uncouplers facilitate oxidation of substrates without ATP generation.

THANK YOU