

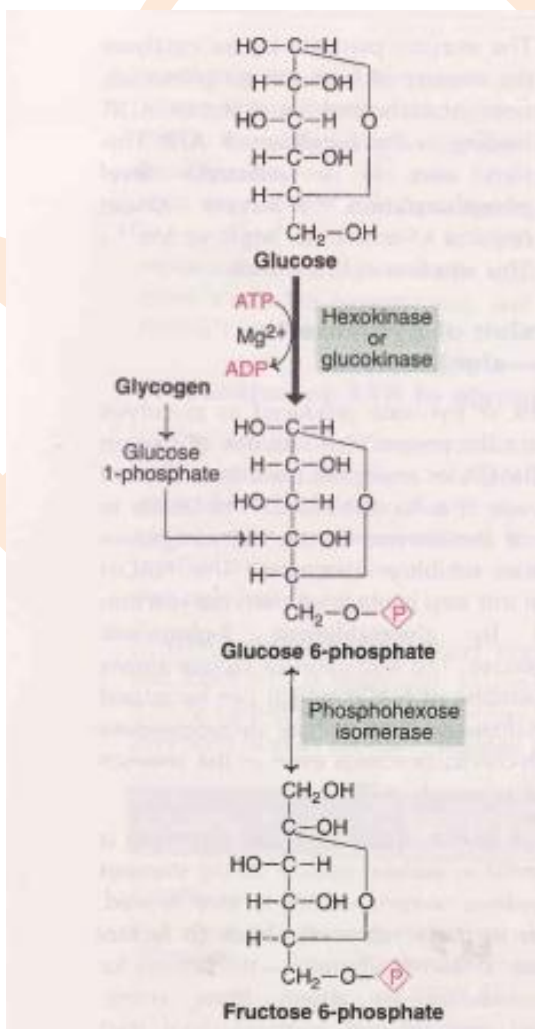
## BIOCHEMISTRY- BP203T

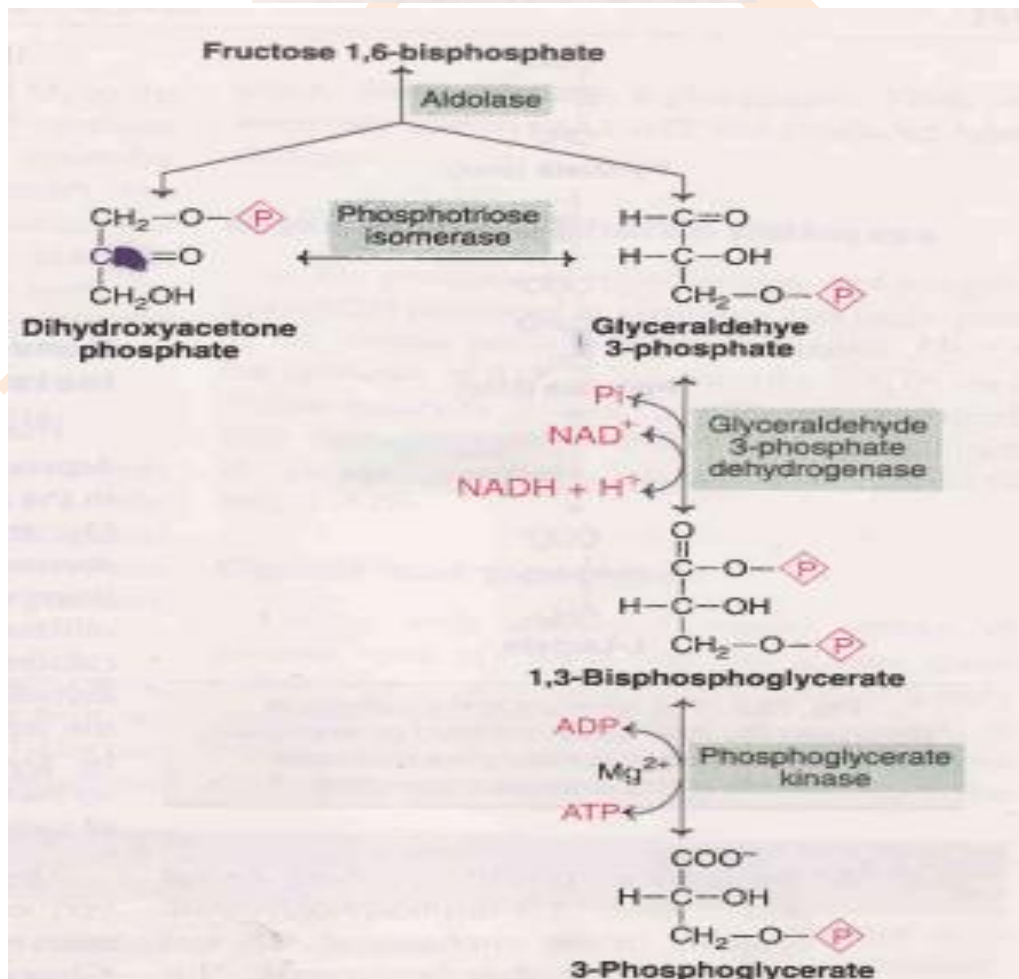
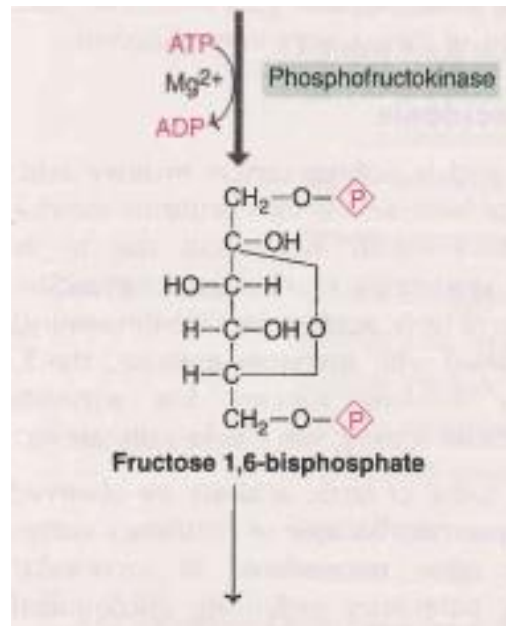
UNIT: 1 CARBHOHYDRATES

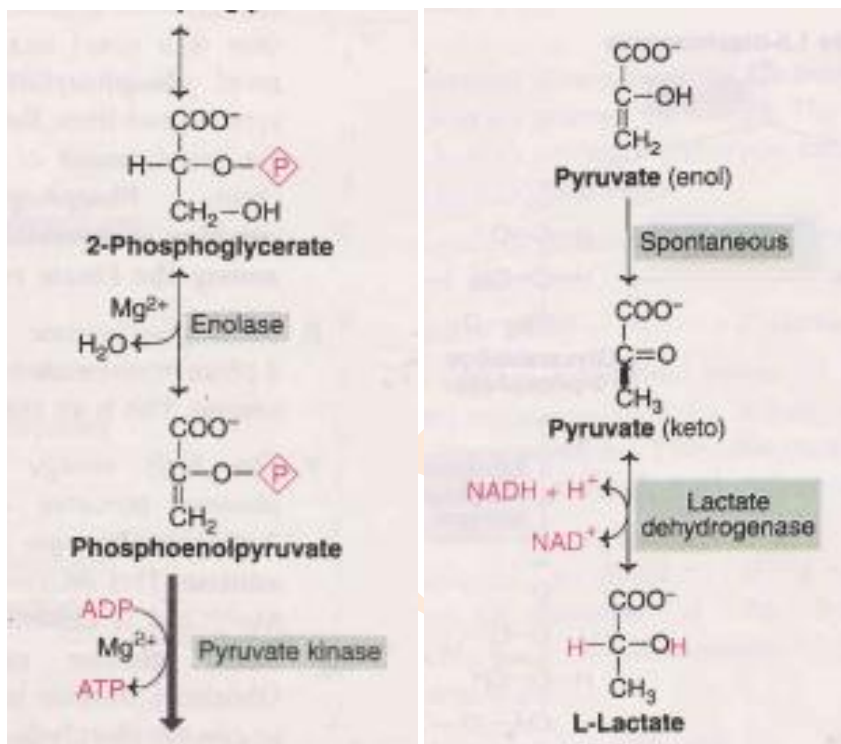
CLASS:2

TOPIC: GLYCOLYSIS

GLYCOLYSIS







The pathway can be divided into three distinct phases:

- A. Energy investment phase or priming stage
- B. Splitting phase
- C. Energy generation phase

I. Glucose is phosphorylated to glucose 6-phosphate by hexokinase or glucokinase (both are isoenzymes). This is an irreversible reaction, dependent on ATP and  $Mg^{2+}$ .

- The enzyme hexokinase is present in almost all the tissues. It catalyses the phosphorylation of various hexoses (fructose, mannose etc.), has low  $K_m$  for substrates and is inhibited by glucose 6-phosphate.
- Glucose-6-phosphate is impermeable to the cell membrane. It is a central molecule with a variety of metabolic fates- glycolysis, glycogenesis, gluconeogenesis and pentose phosphate pathway
- II. Glucose-6-phosphate undergoes isomerization to give fructose-6-phosphate in the presence of the enzyme phospho-hexose isomerase and  $Mg^{2+}$ .
- III. Fructose-6-phosphate is phosphorylated to fructose-1,6-bisphosphate by phosphofructokinase (PFK). This is an irreversible and regulatory step in glycolysis.

VI. Glyceraldehyde-3-phosphate dehydrogenase converts glyceraldehyde 3-phosphate to 1,3-bisphosphoglycerate.

- This step is important as it is involved in the formation of  $\text{NADH} + \text{H}^+$  and a high-energy compound, 1,3-bisphosphoglycerate.
- In aerobic conditions, NADH passes through the electron transport chain and 6 ATP ( $2 \times 3 \text{ ATP}$ ) are synthesized by oxidative phosphorylation.

VII. The enzyme phosphoglycerate kinase acts on 1,3-bisphosphoglycerate, resulting in the synthesis of ATP and the formation of 3-phosphoglycerate.

- This step is a good example of substrate-level phosphorylation, since ATP is synthesized from the substrate without the involvement of the electron transport chain.
- VIII. 3-Phosphoglycerate is converted to 2-phosphoglycerate by phosphoglycerate mutase. This is an isomerization reaction.
- IX. The high-energy compound phosphoenol pyruvate is generated from 2-phosphoglycerate by the enzyme enolase. This enzyme requires  $\text{Mg}^{2+}$  or  $\text{Mn}^{2+}$ .
- X The enzyme pyruvate kinase catalyses the transfer of high-energy phosphate from phosphoenol pyruvate to ADP, leading to the formation of ATP. This step is also a substrate-level phosphorylation.

#### Generation of ATP in glycolysis

| Pathway    | Enzyme (method of ATP synthesis)                                                      | Number of ATP synthesized |
|------------|---------------------------------------------------------------------------------------|---------------------------|
| Glycolysis | Glyceraldehyde 3-phosphate dehydrogenase<br>(2 NADH, ETC, oxidative phosphorylation)  | 6*                        |
|            | Phosphoglycerate kinase (substrate level phosphorylation)                             | 2                         |
|            | Pyruvate kinase (substrate level phosphorylation)                                     | 2                         |
|            | Two ATP are consumed in the reactions catalysed by hexokinase and phosphofructokinase | -2                        |
|            | <b>Net ATP synthesis in glycolysis in aerobic condition</b>                           | <b>8</b>                  |