

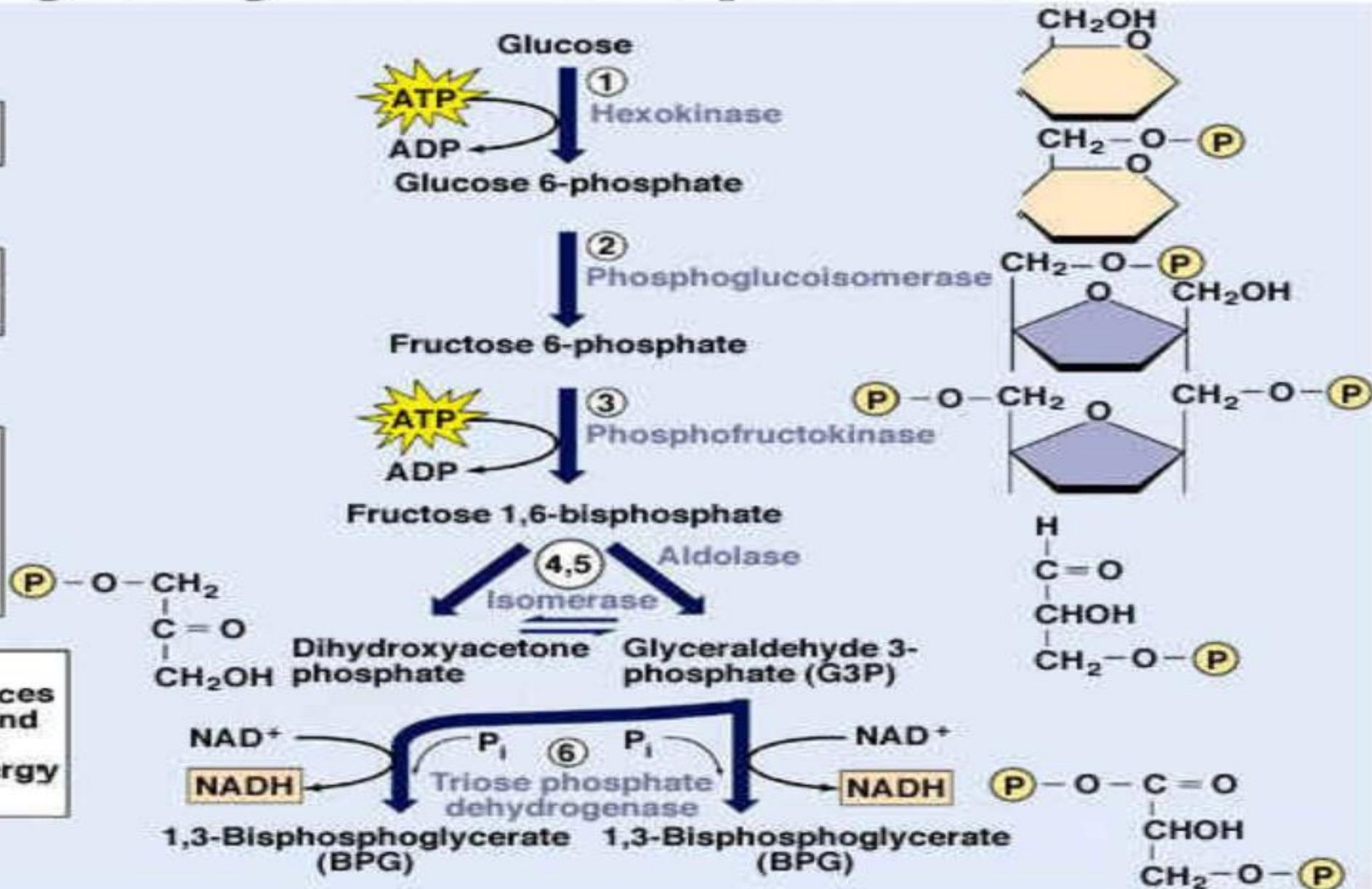
# Glycolysis — Steps 1-6

1. Phosphorylation of glucose by ATP

2-3. Rearrangement, followed by a second ATP phosphorylation.

4-5. The six-carbon molecule is split into two three-carbon molecules—one G3P, another that is converted into G3P in another reaction.

6. Oxidation followed by phosphorylation produces two NADH molecules and two molecules of BPG, each with one high-energy phosphate bond.



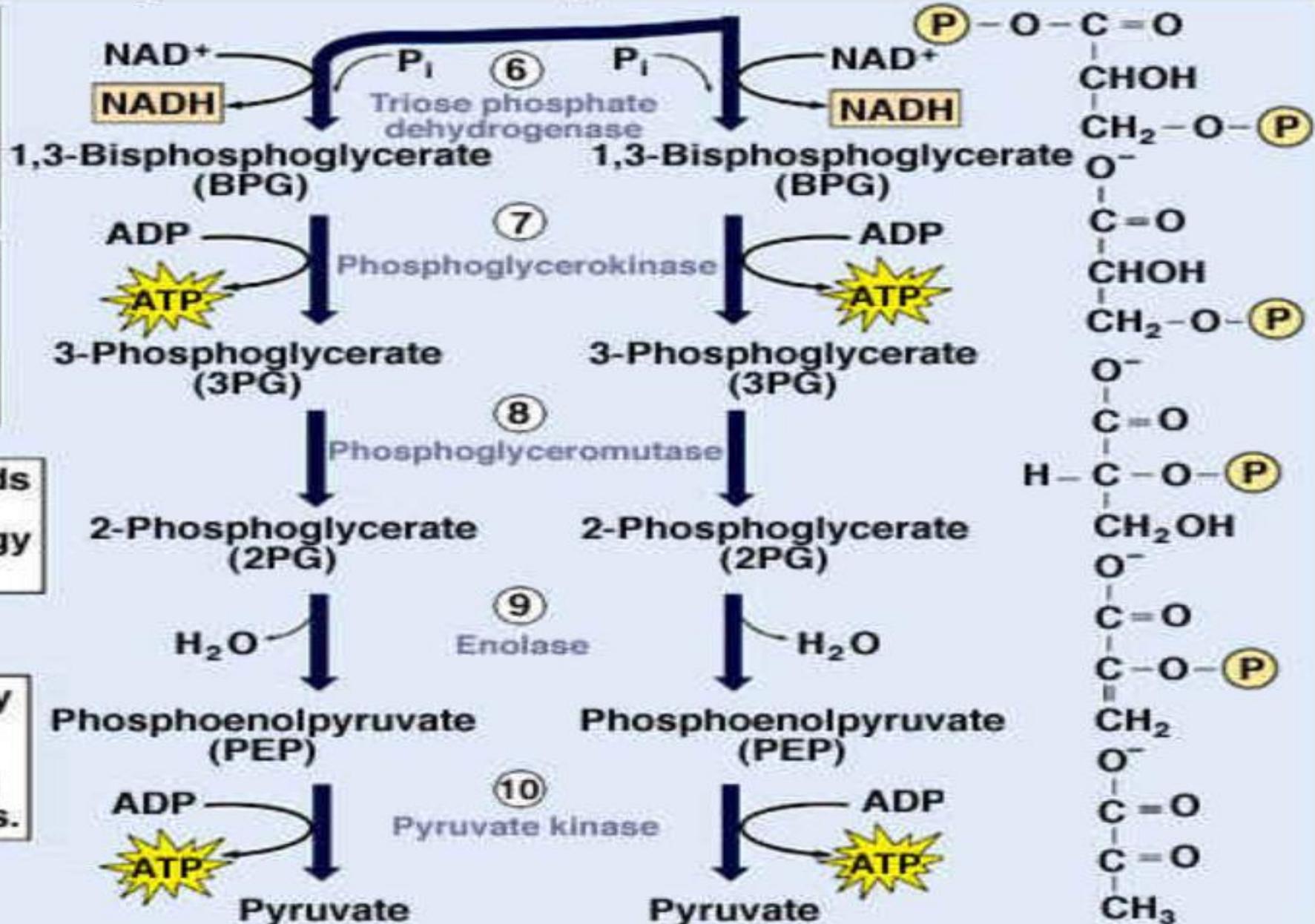
# Glycolysis—Steps 6 – 10

6. Oxidation followed by phosphorylation produces two NADH molecules and two molecules of BPG, each with one high-energy phosphate bond.

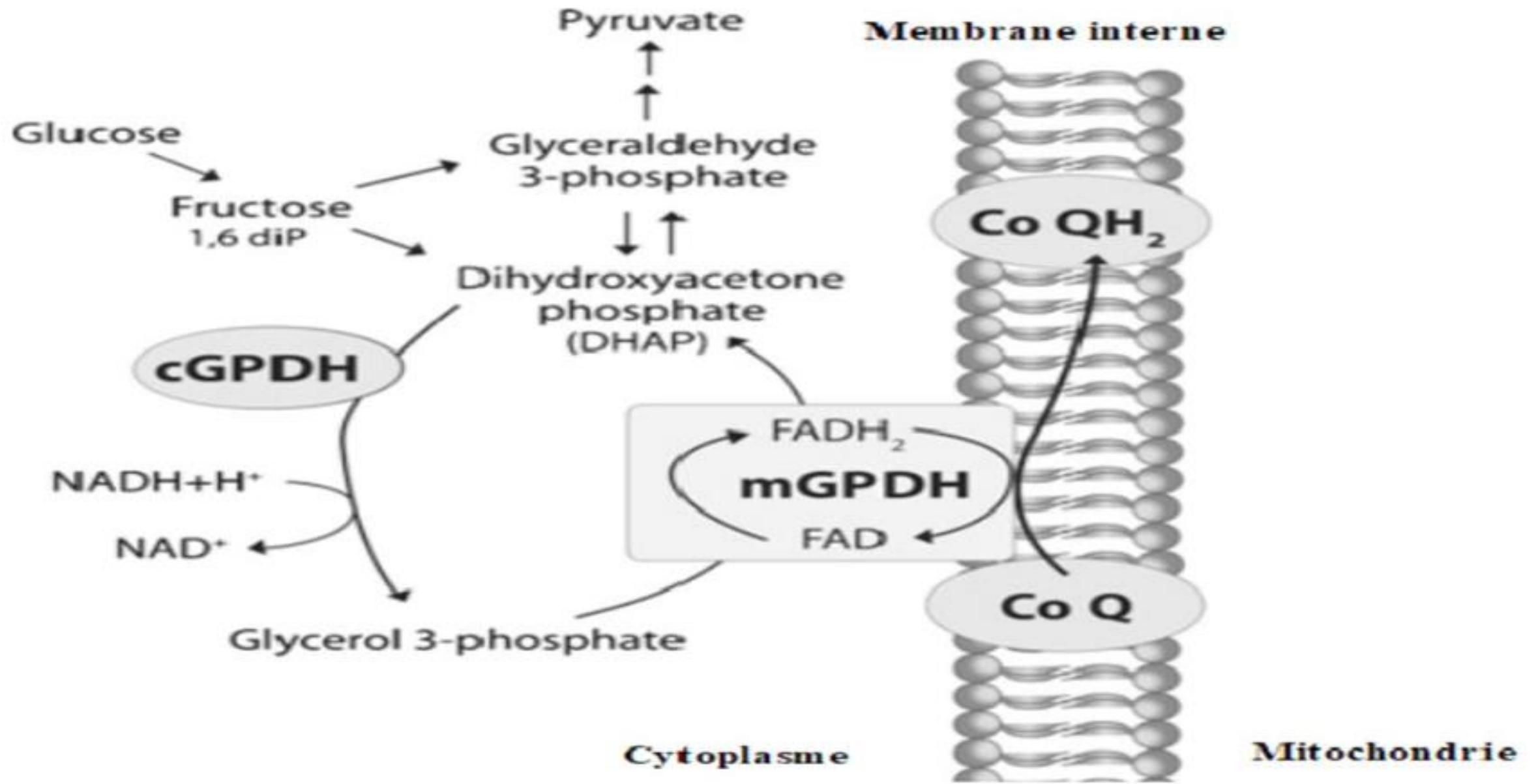
7. Removal of high-energy phosphate by two ADP molecules produces two ATP molecules and leaves two 3PG molecules.

8-9. Removal of water yields two PEP molecules, each with a high-energy phosphate bond.

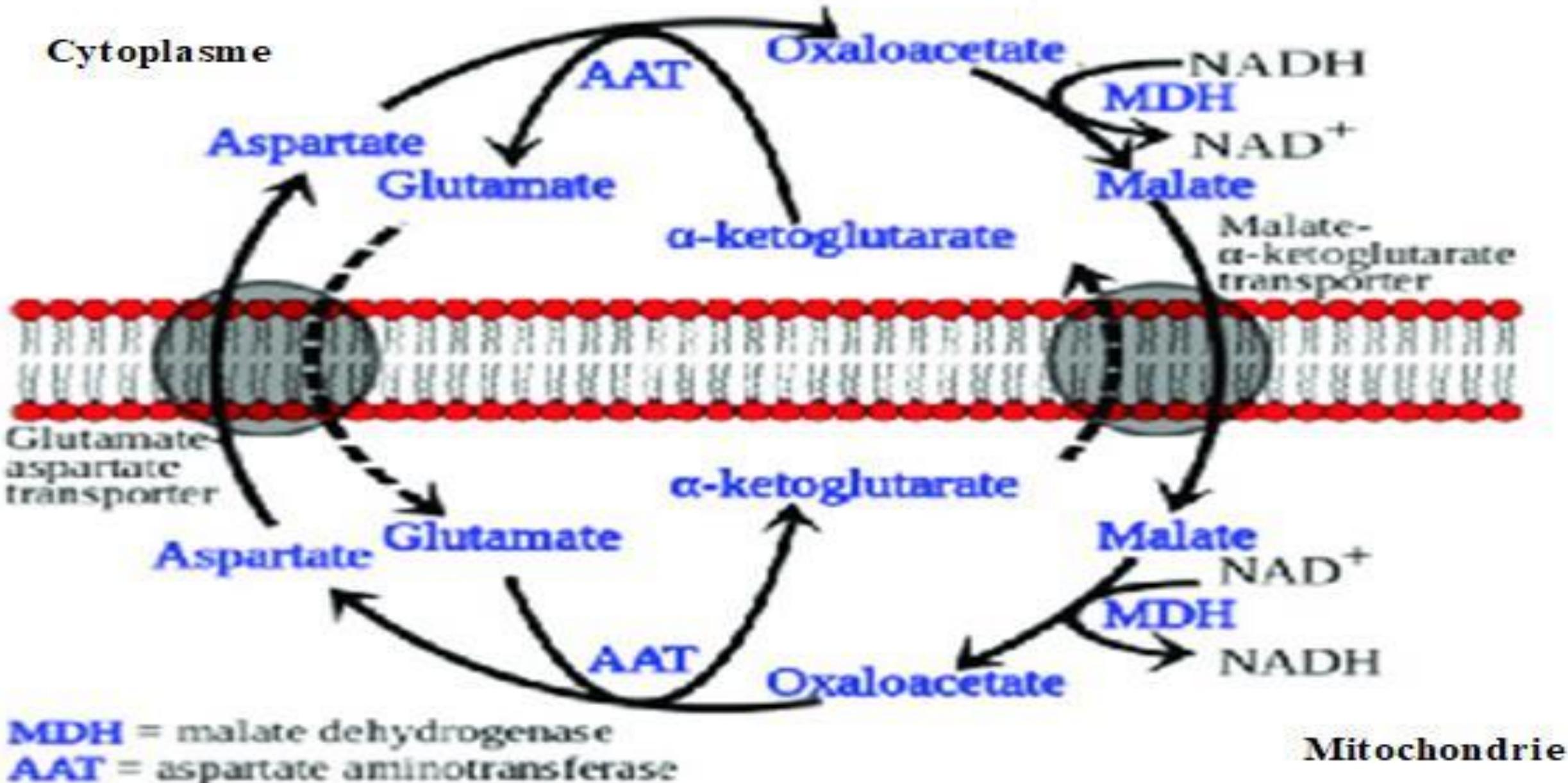
10. Removal of high-energy phosphate by two ADP molecules produces two ATP molecules and two pyruvate molecules.



# Glycerol 3 phosphate shuttle

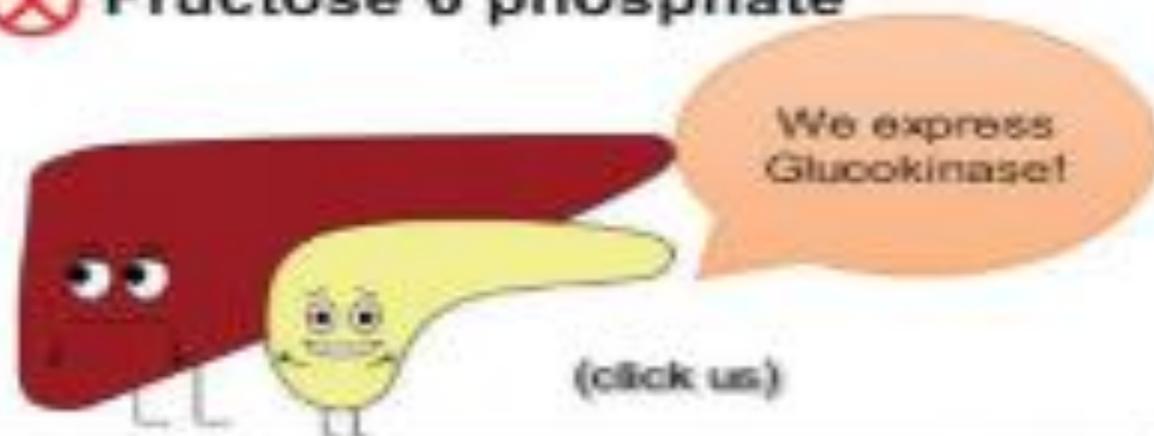


# Aspartate-malate shuttle

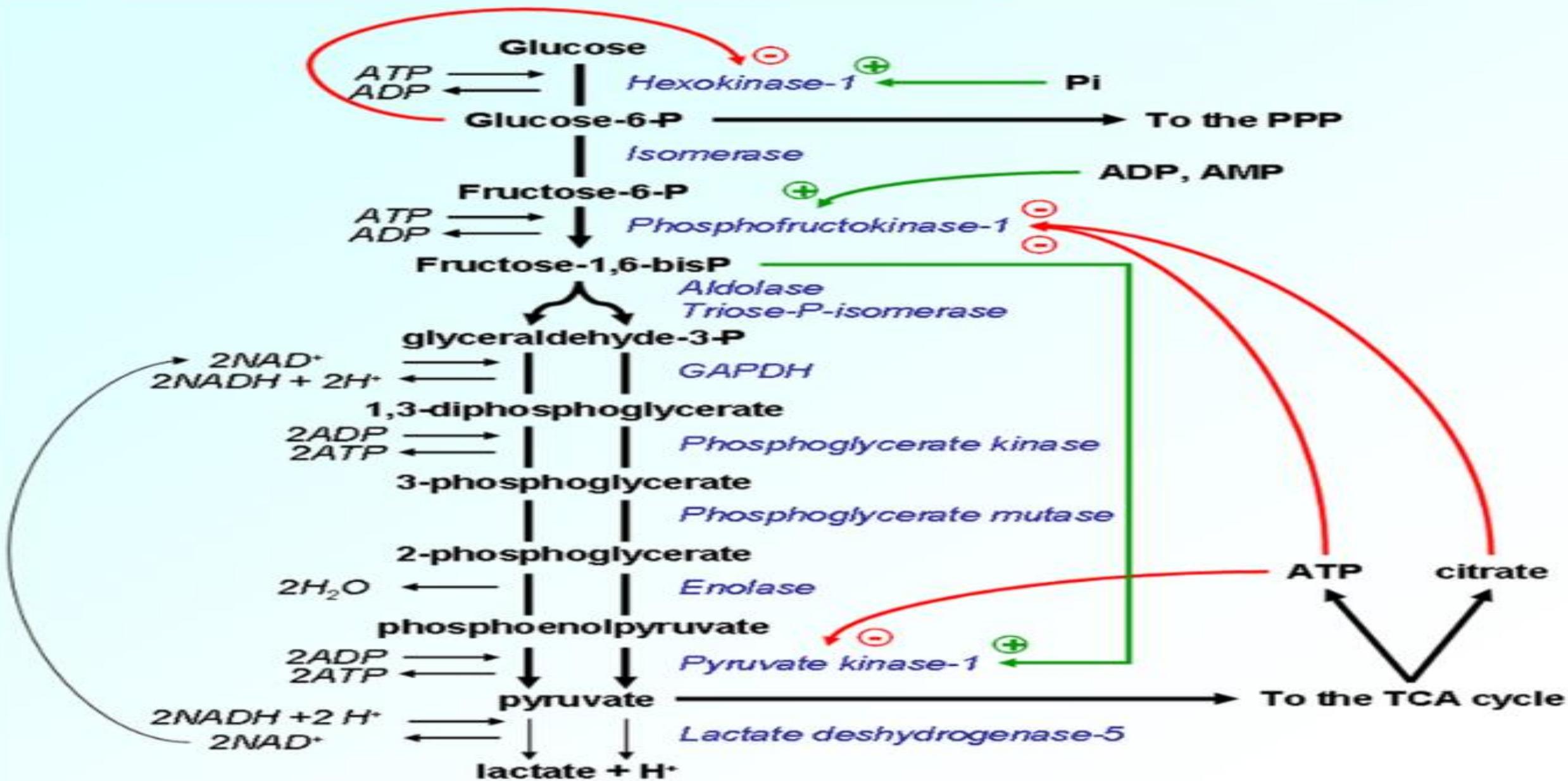


# Hexokinase vs. Glucokinase

- |   |  |
|---|--|
| <ul style="list-style-type: none"><li>• Expressed in most tissues</li><li>• High Affinity (Low <math>K_m</math>)</li><li>• Low capacity (Low <math>V_{max}</math>)</li><li>• NOT induced by insulin</li><li>• Inhibited by  <b>Glucose-6-Phosphate</b></li></ul> | <ul style="list-style-type: none"><li>• Expressed in Liver and Beta cells of pancreas</li><li>• Lower affinity (High <math>K_m</math>)</li><li>• High Capacity (High <math>V_{max}</math>)</li><li>• Induced by insulin</li><li>• Inhibited by  <b>Fructose 6 phosphate</b></li></ul> |
|---|--|



# Allosteric regulation of glycolysis enzymes



# Phosphofructokinase 1 regulation

↓ cAMP leads to dephosphorylation of bifunctional enzyme

↑ cAMP leads to phosphorylation of bifunctional enzyme

Insulin

Glucagon

+

+

↑ Fructose 2,6-bisphosphate

phospho fructokinase-2

Fructose 6-phosphate

fructose 2,6-bisphosphatase

Fructose 2,6-bisphosphate ↓

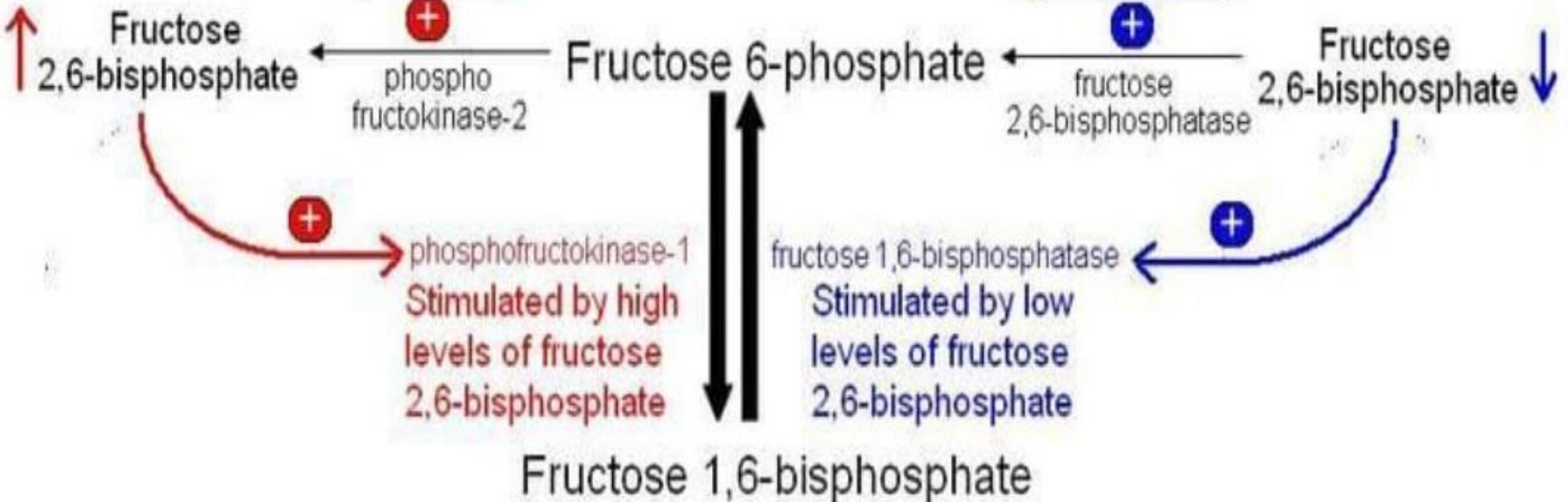
+

+

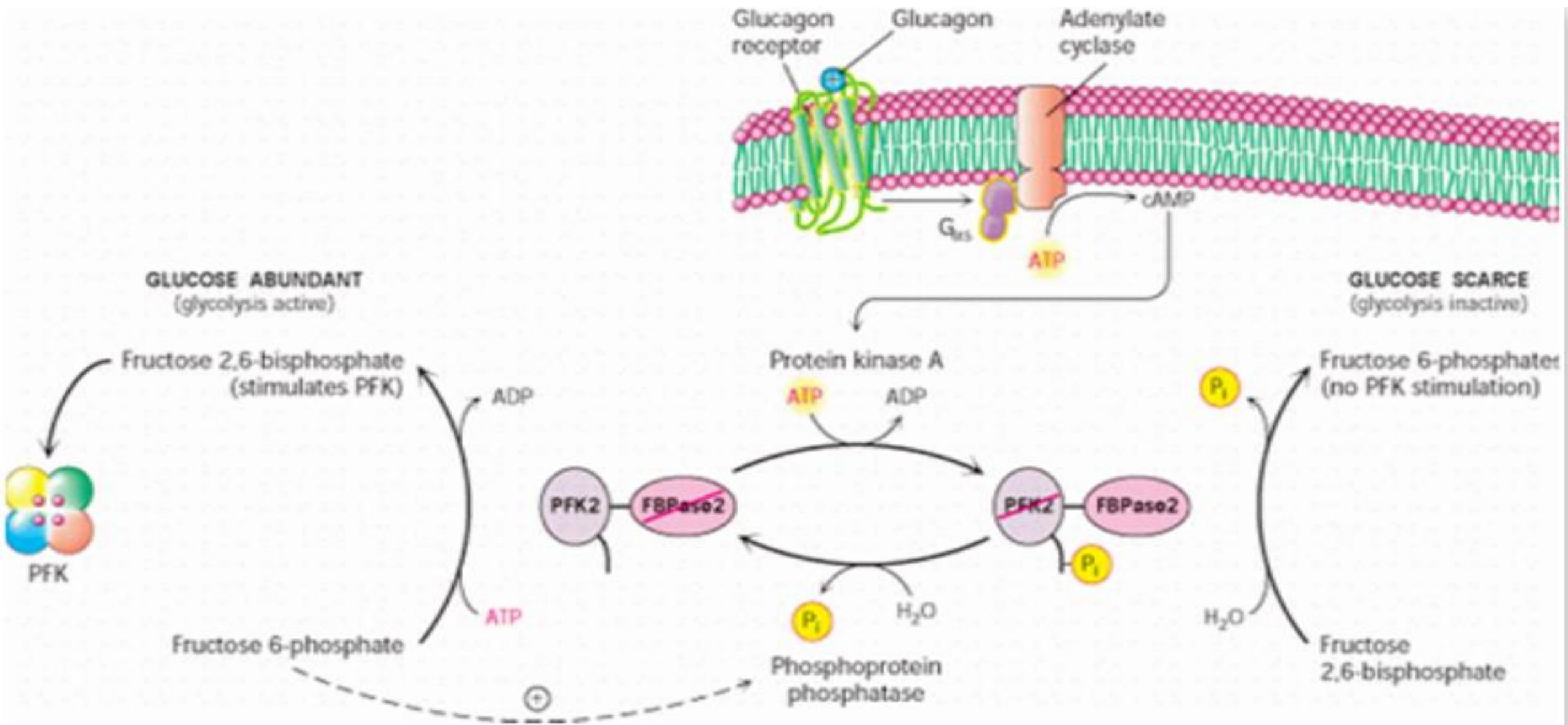
phosphofructokinase-1  
Stimulated by high levels of fructose 2,6-bisphosphate

fructose 1,6-bisphosphatase  
Stimulated by low levels of fructose 2,6-bisphosphate

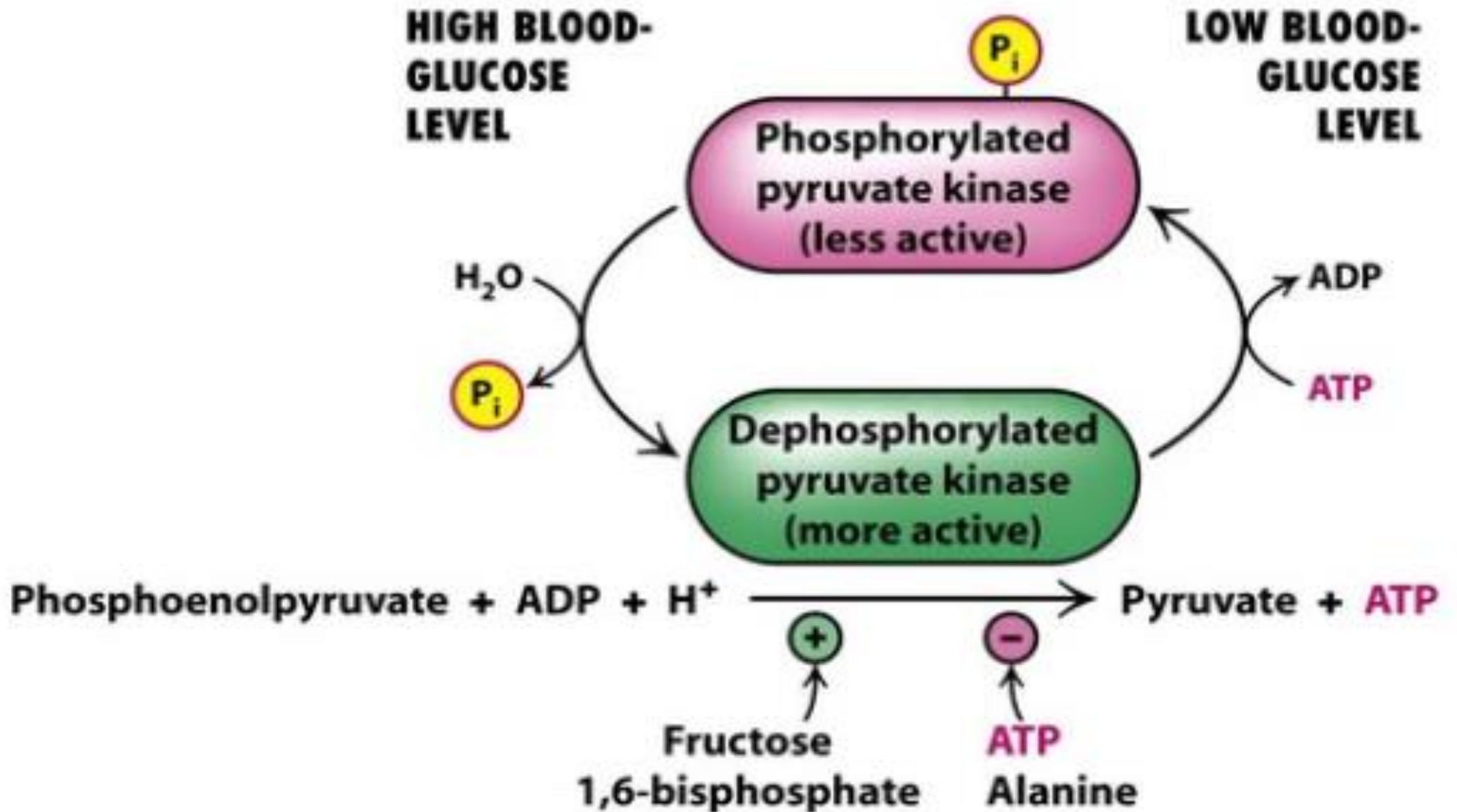
Fructose 1,6-bisphosphate



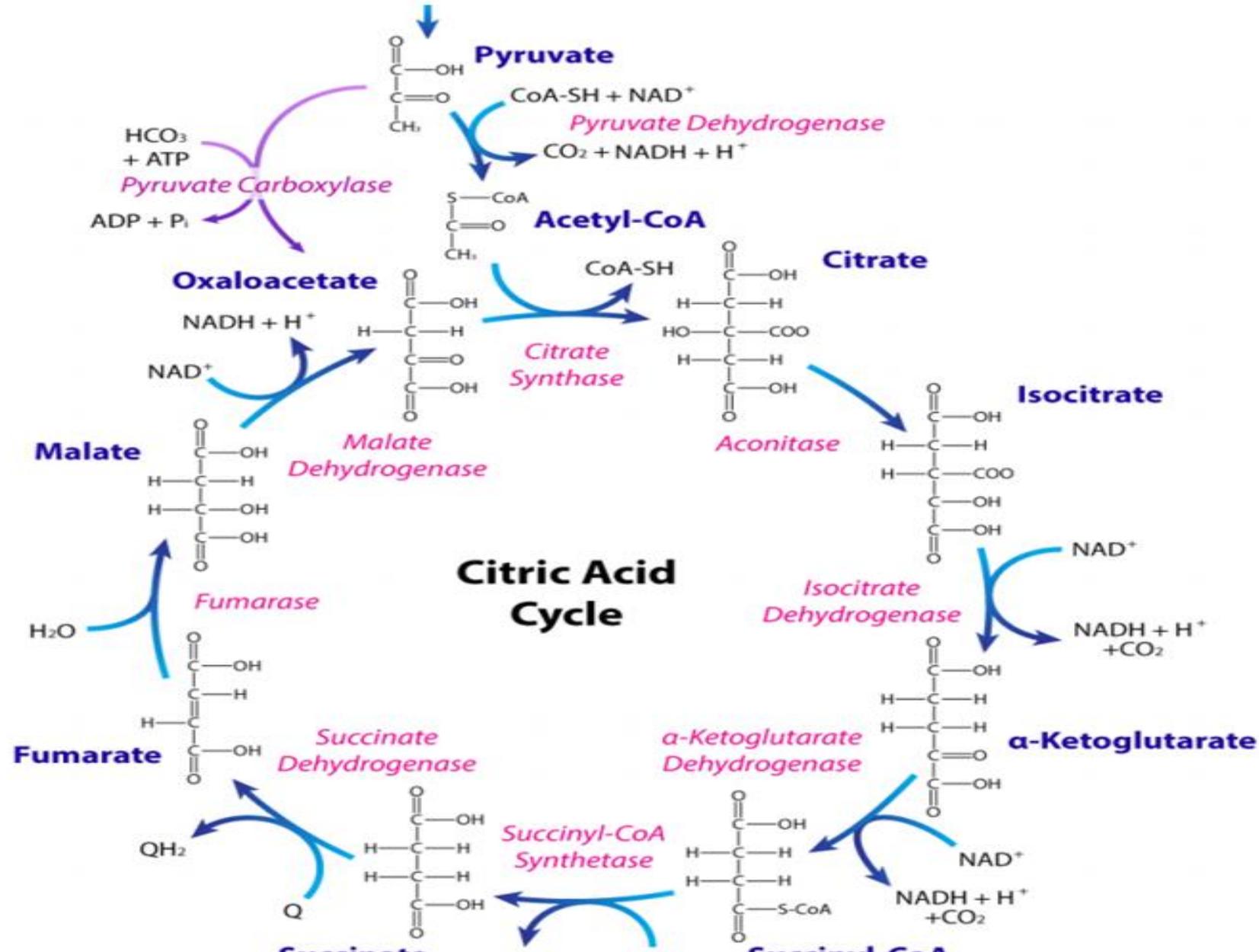
# Phosphofructokinase 1 regulation



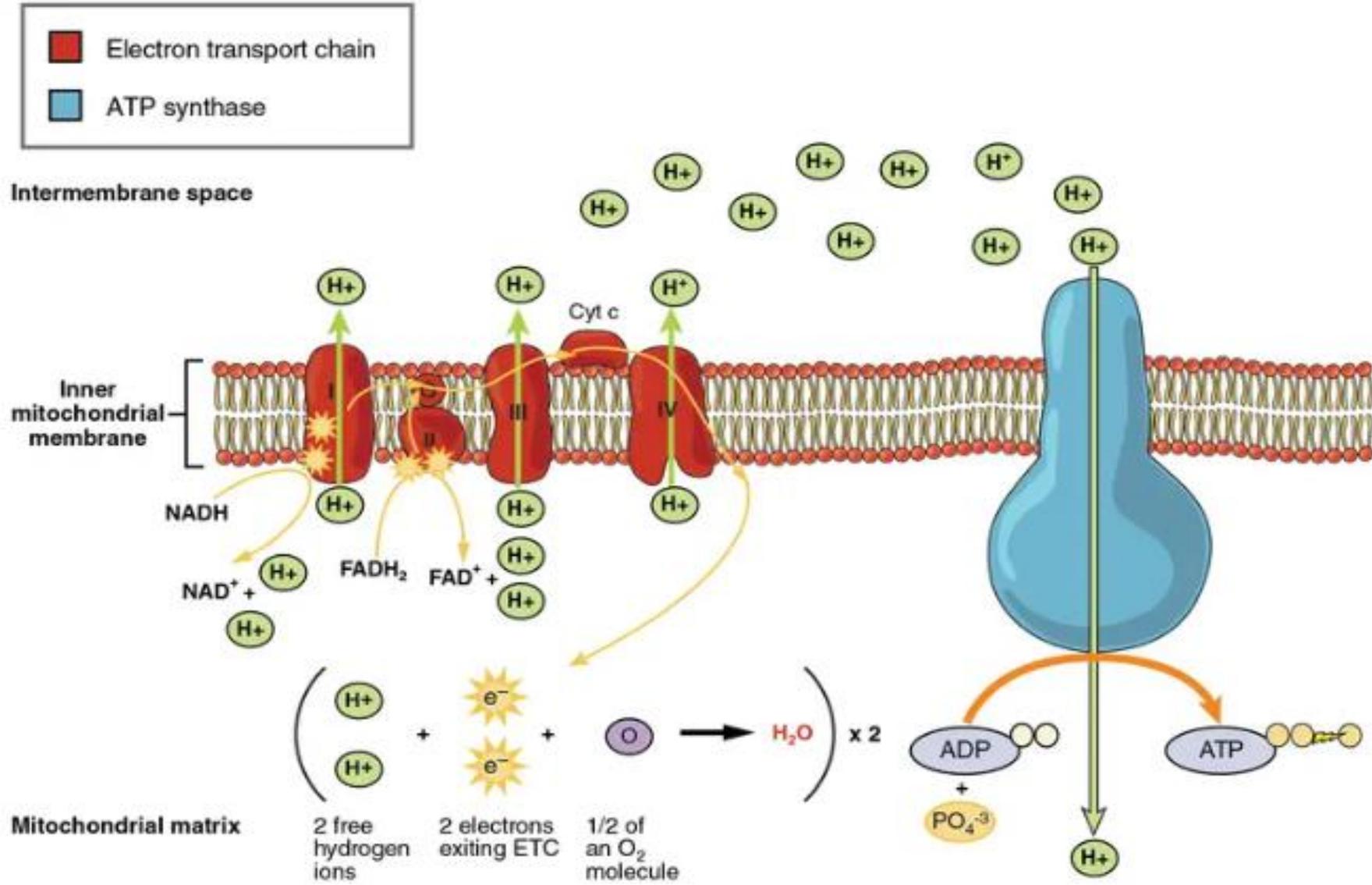
# Pyruvate kinase regulation



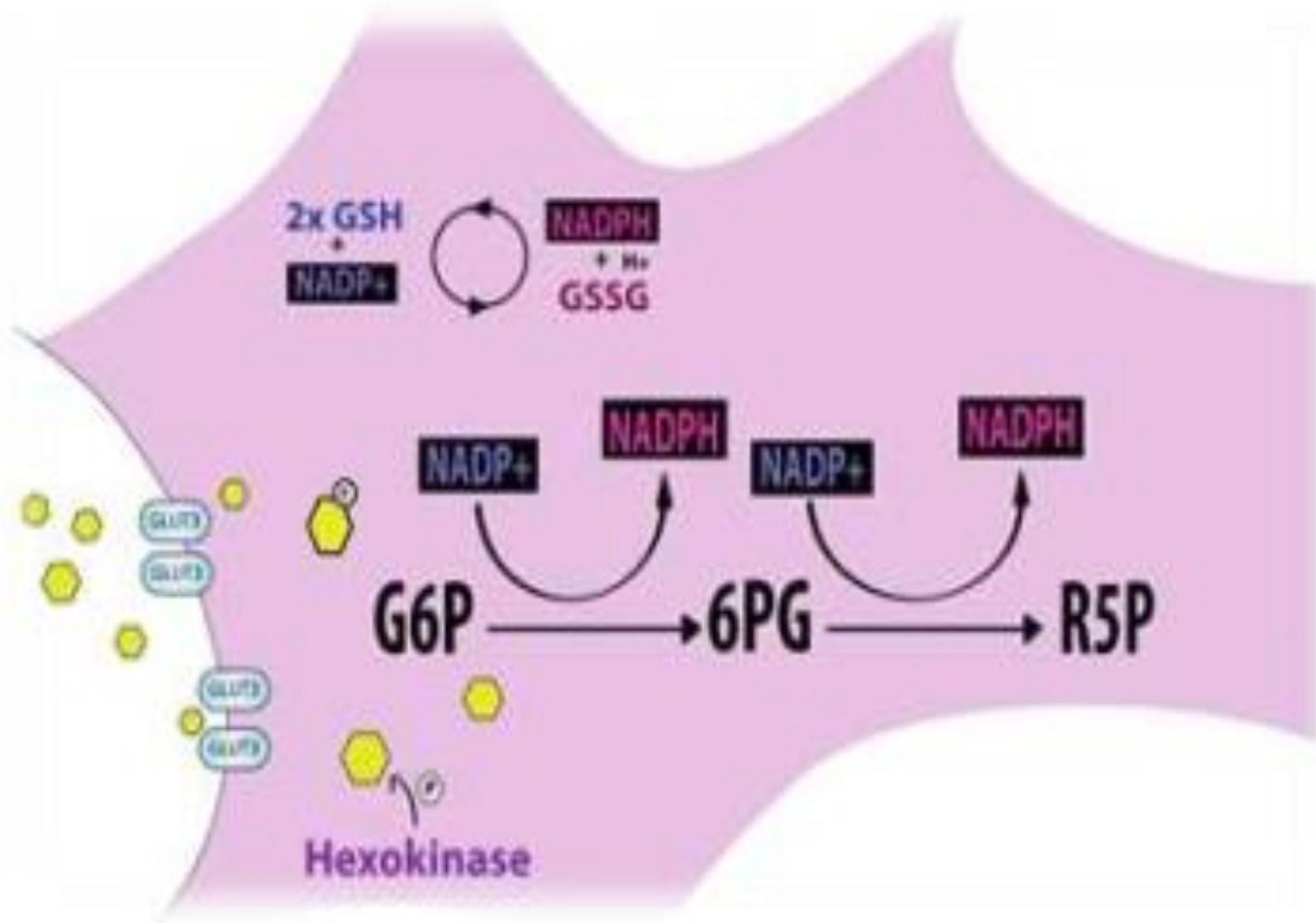
# Citric acid cycle



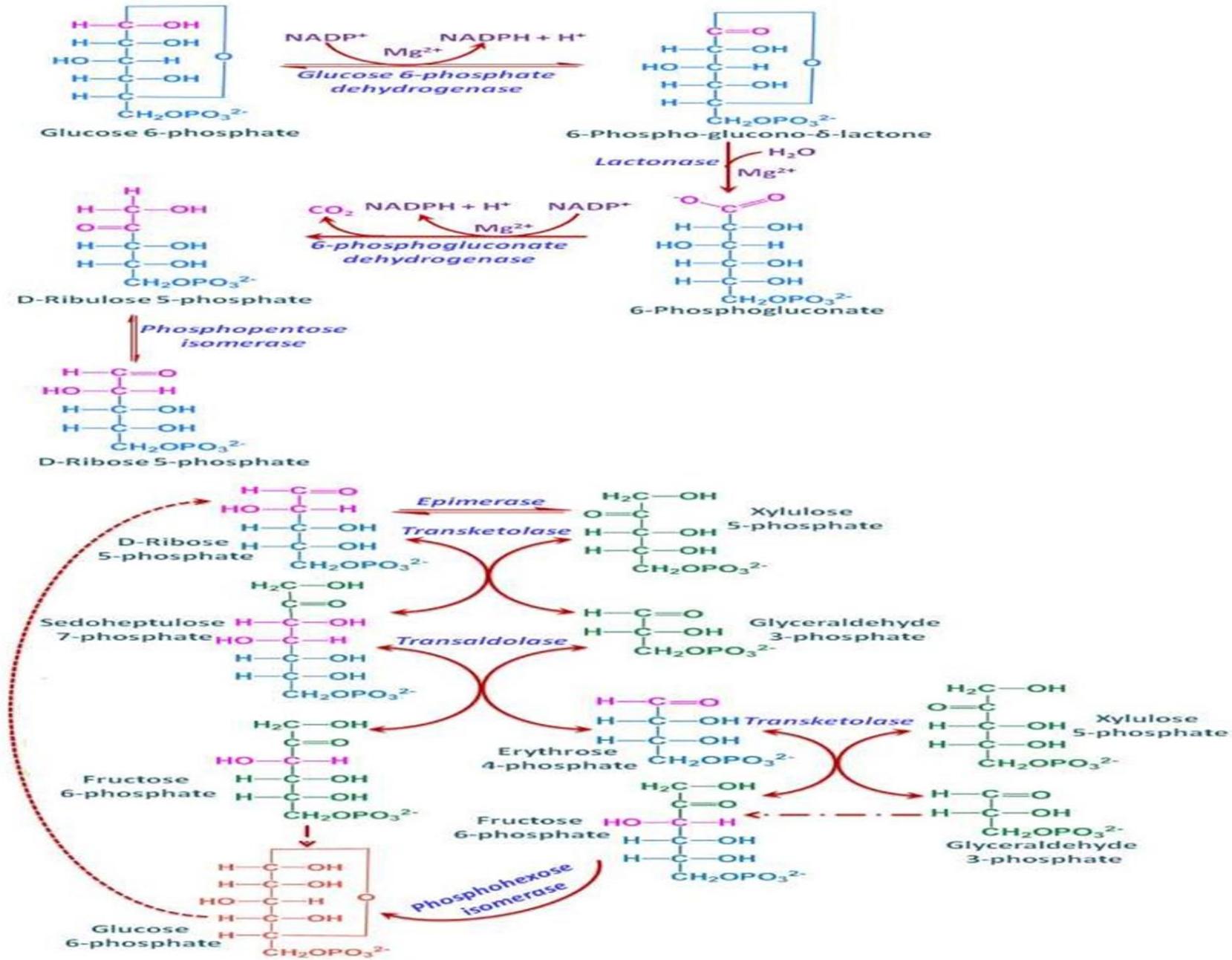
# Oxidative phosphorylation mechanism



# Pentose phosphate pathway

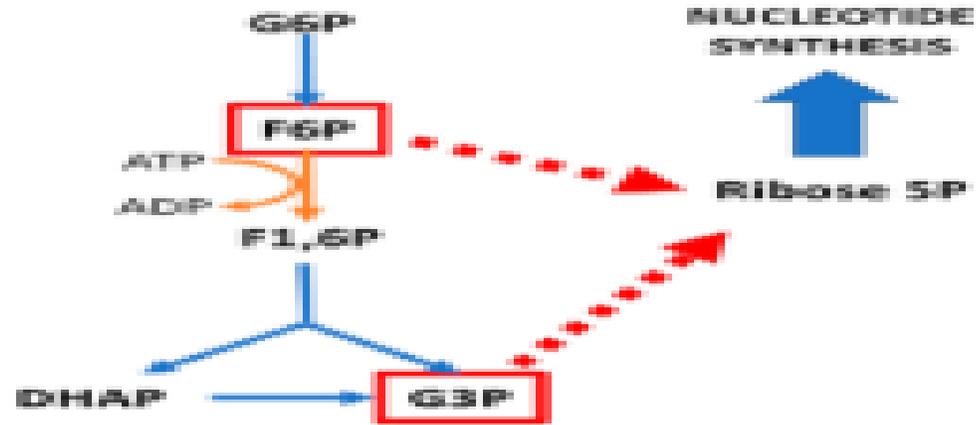


# Pentose phosphate pathway

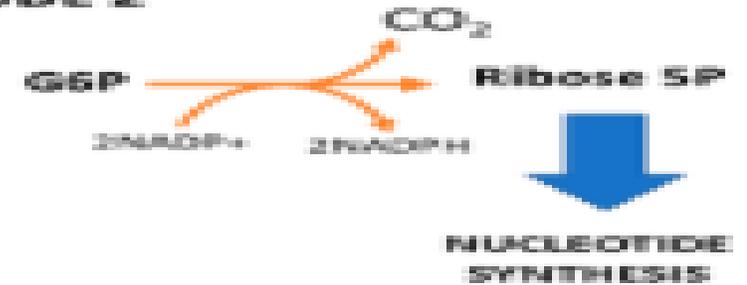


# Pentose phosphate pathway

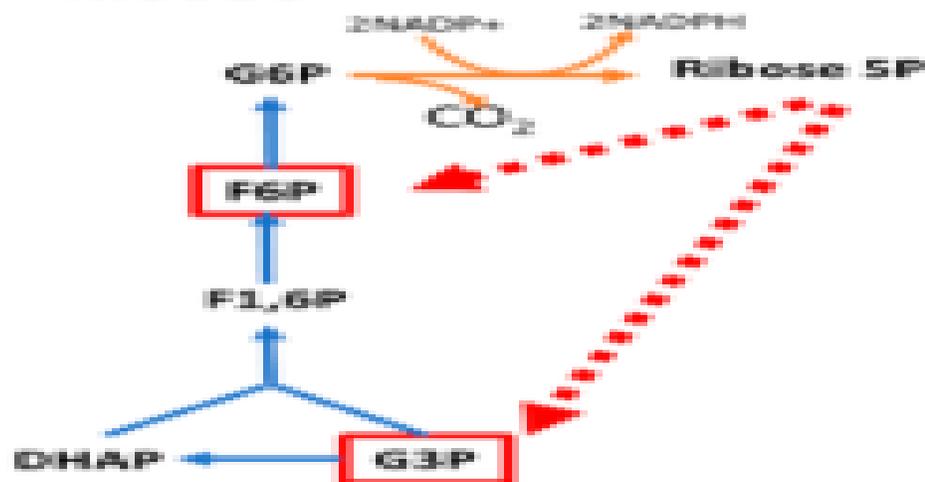
**MODE 1**



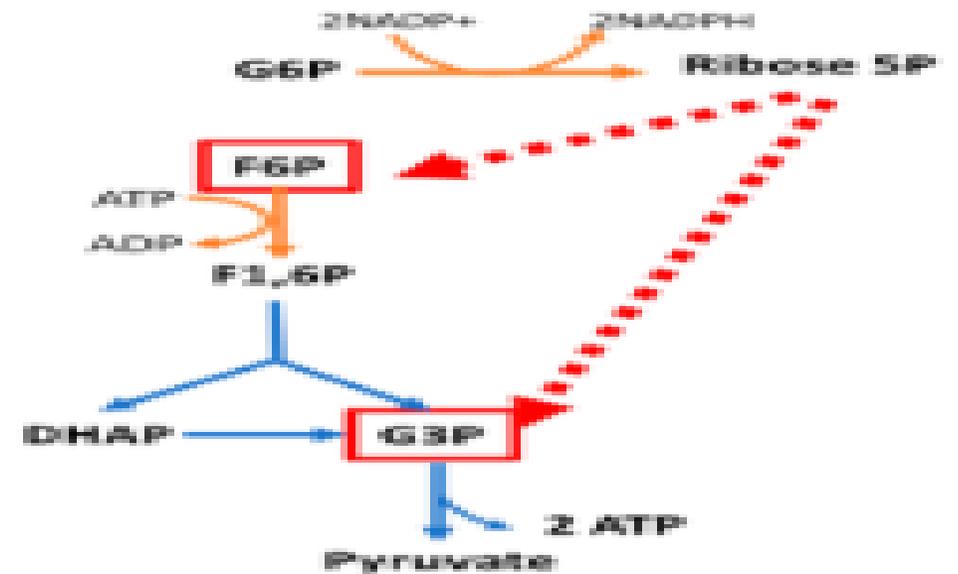
**MODE 2**



**MODE 3**

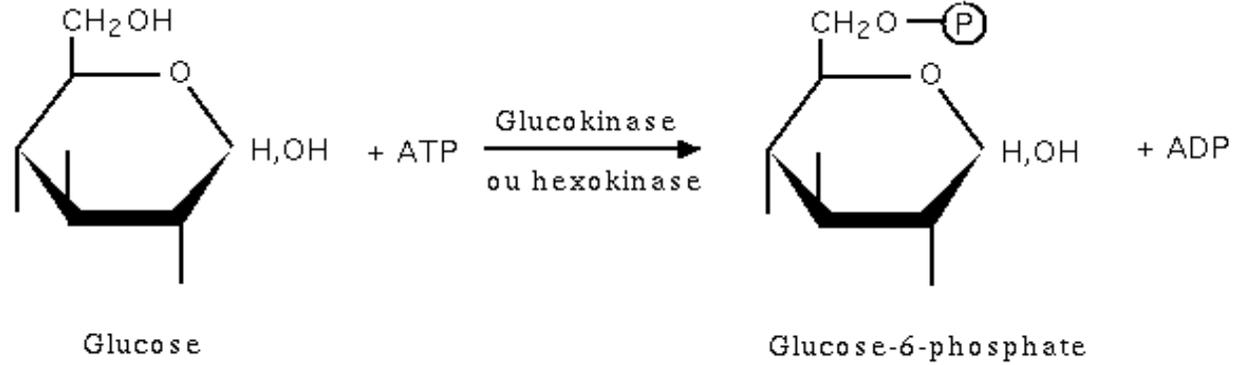


**MODE 4**

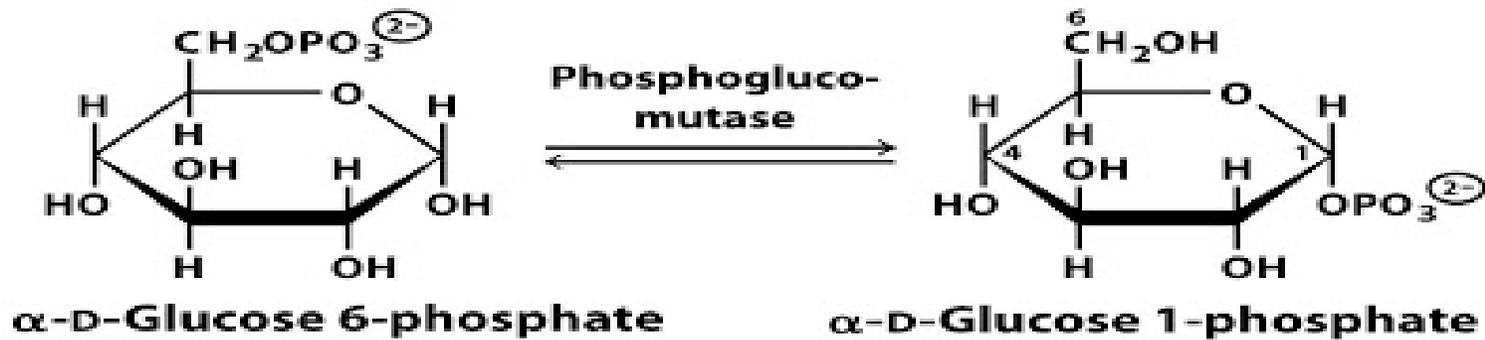




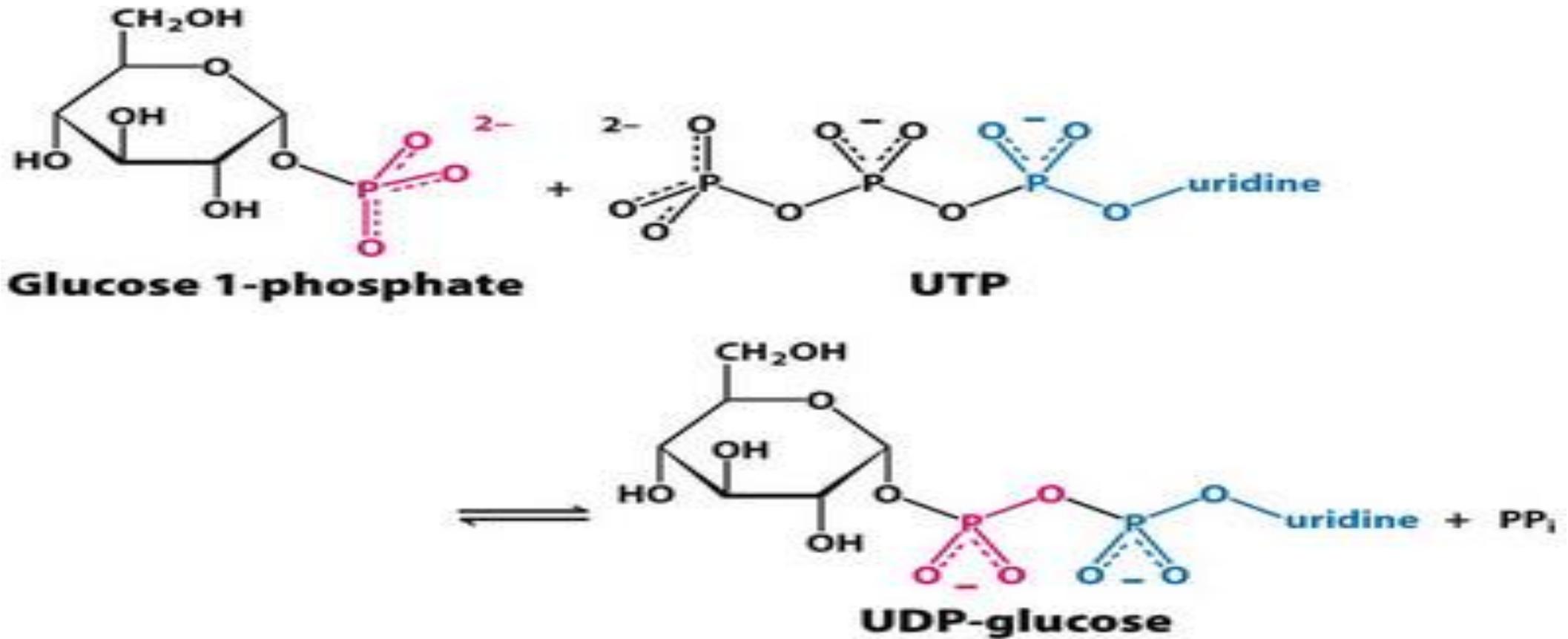
## Reaction 1: Phosphorylation of glucose



## Reaction 2: Conversion of glucose 6-P into glucose 1-P

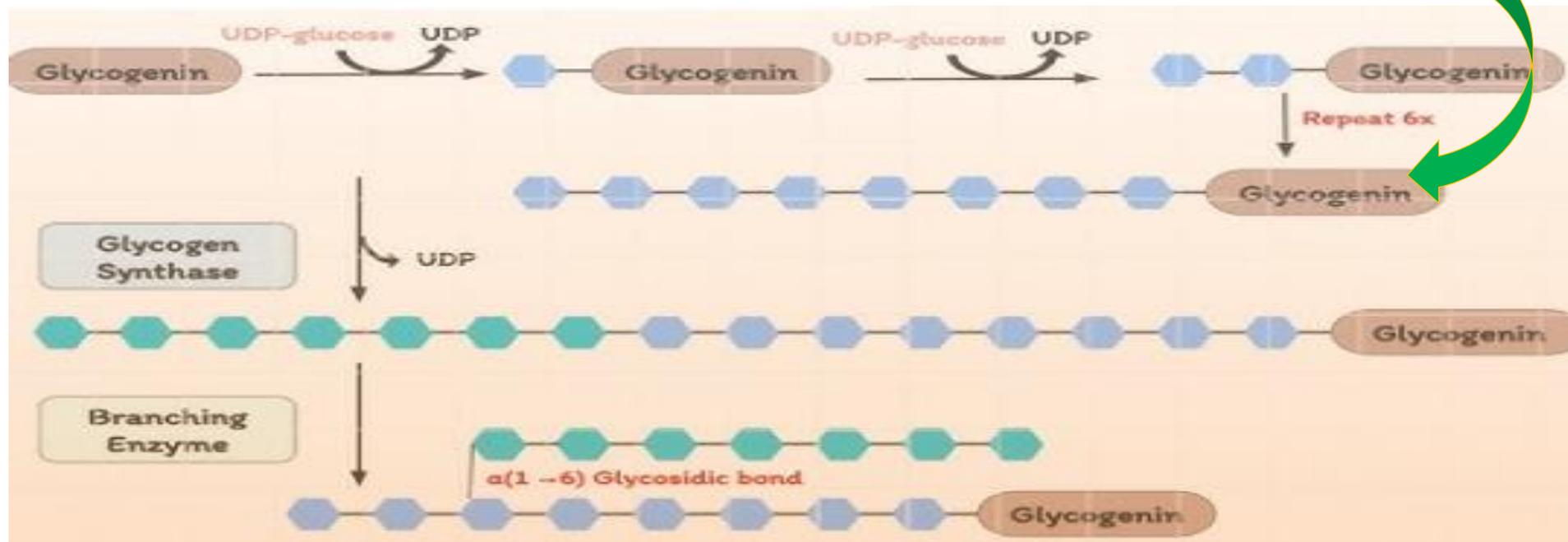
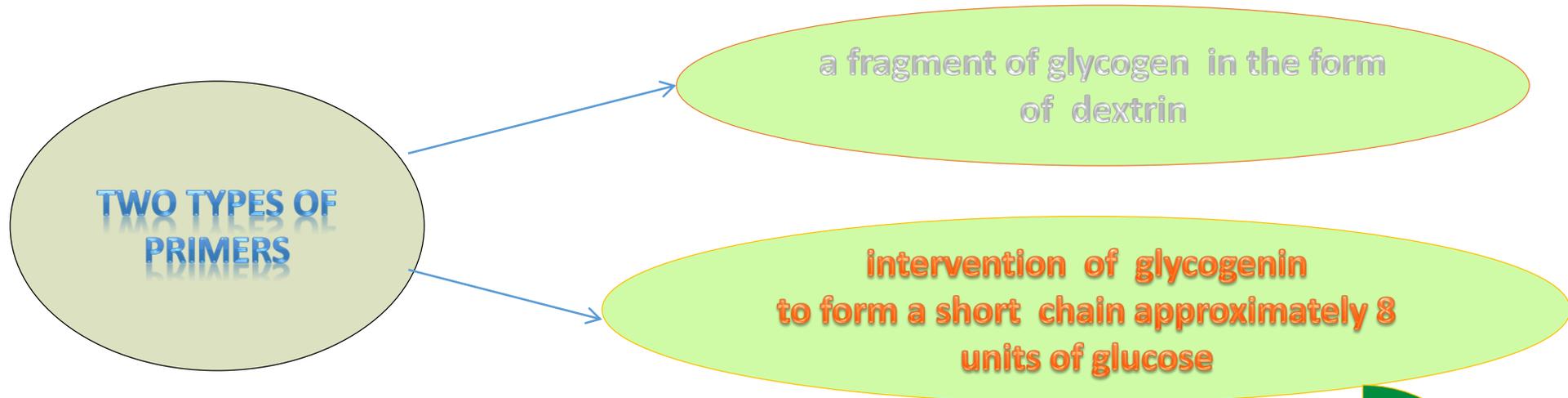


Reaction 3: formation of UDP-glucose by UDP-glucose pyrophosphorylase



# Glycogenesis

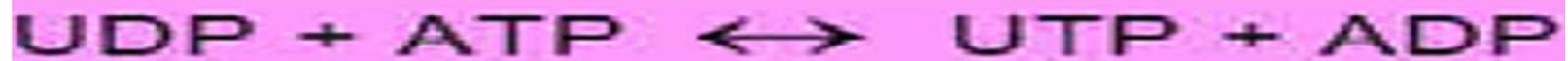
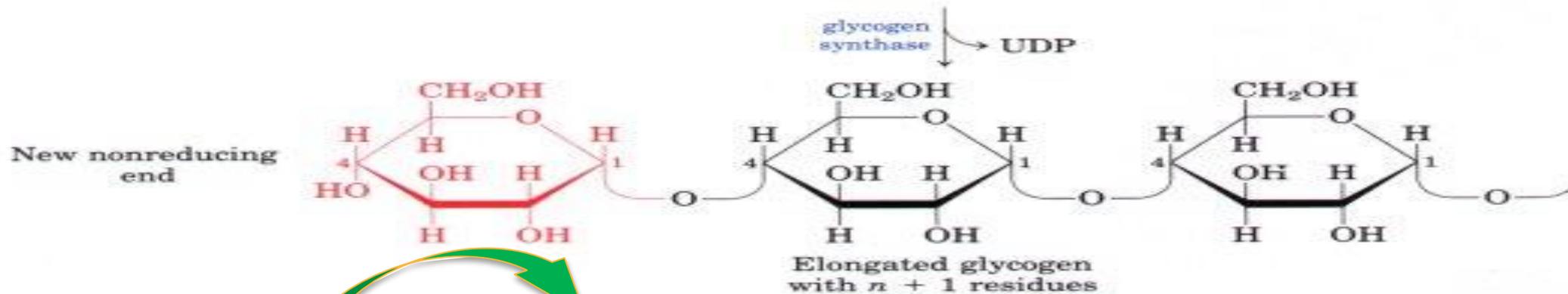
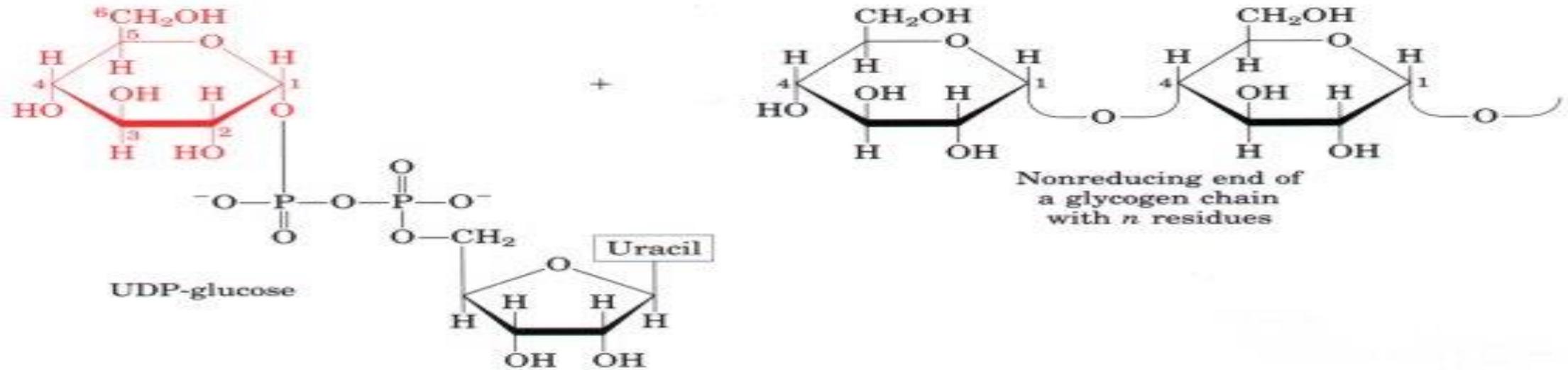
## Reaction 4: **Synthesis** of a primer to initiate the glycogen formation

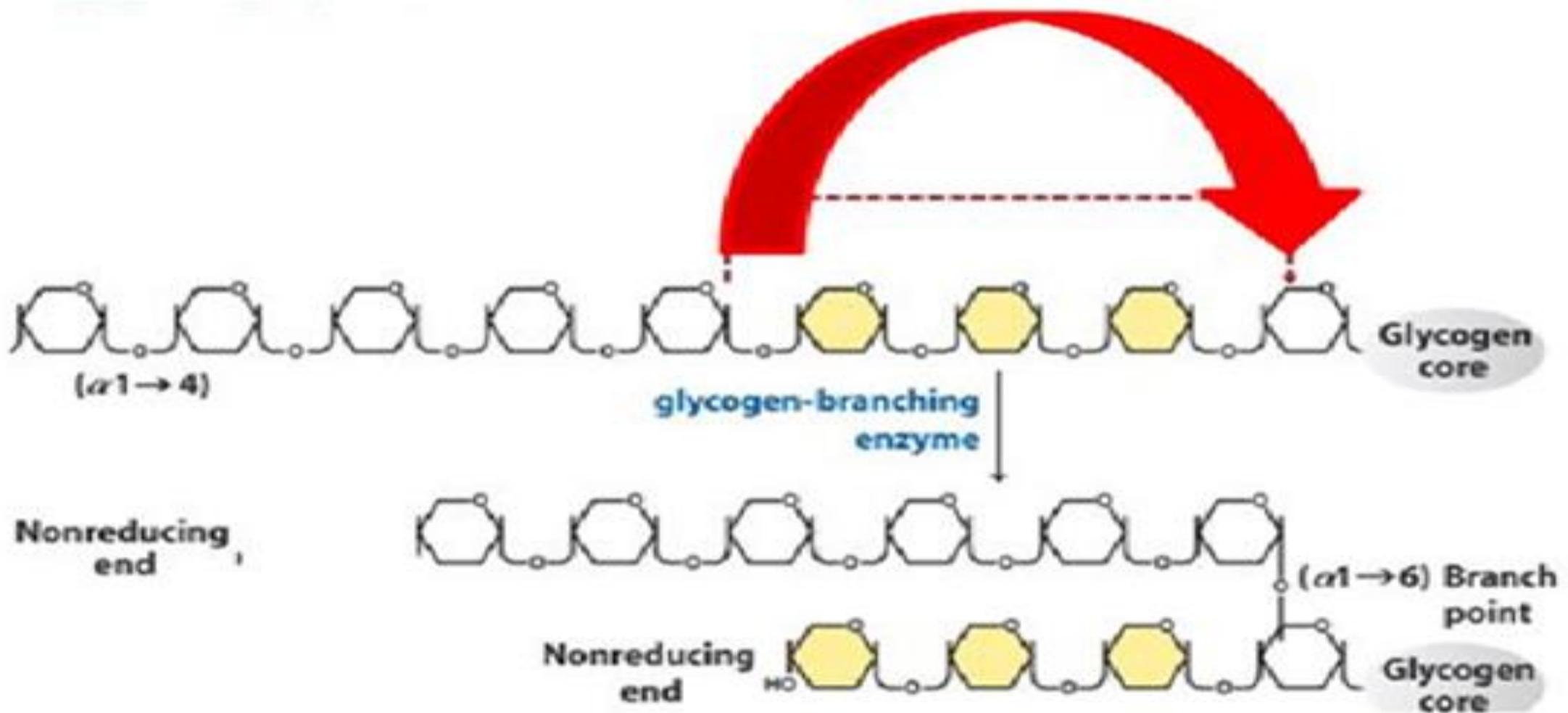


# Glycogenesis

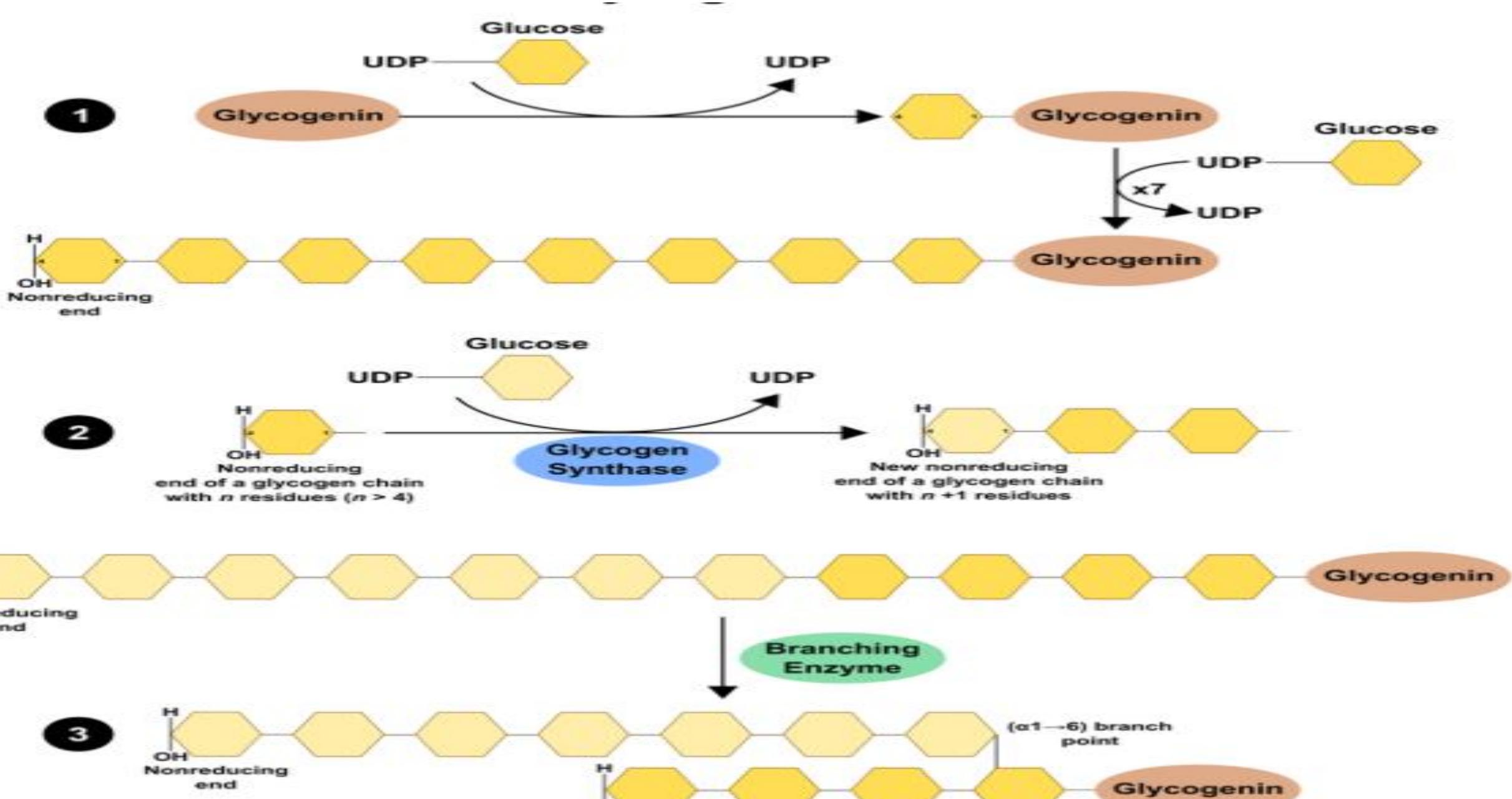
## Reaction 5: Elongation of the chain catalyzed by glycogen synthetase

New glucosyl units added to nonreducing end terminal residues of glycogen and the formation of  $\alpha(1,4)$  linkage

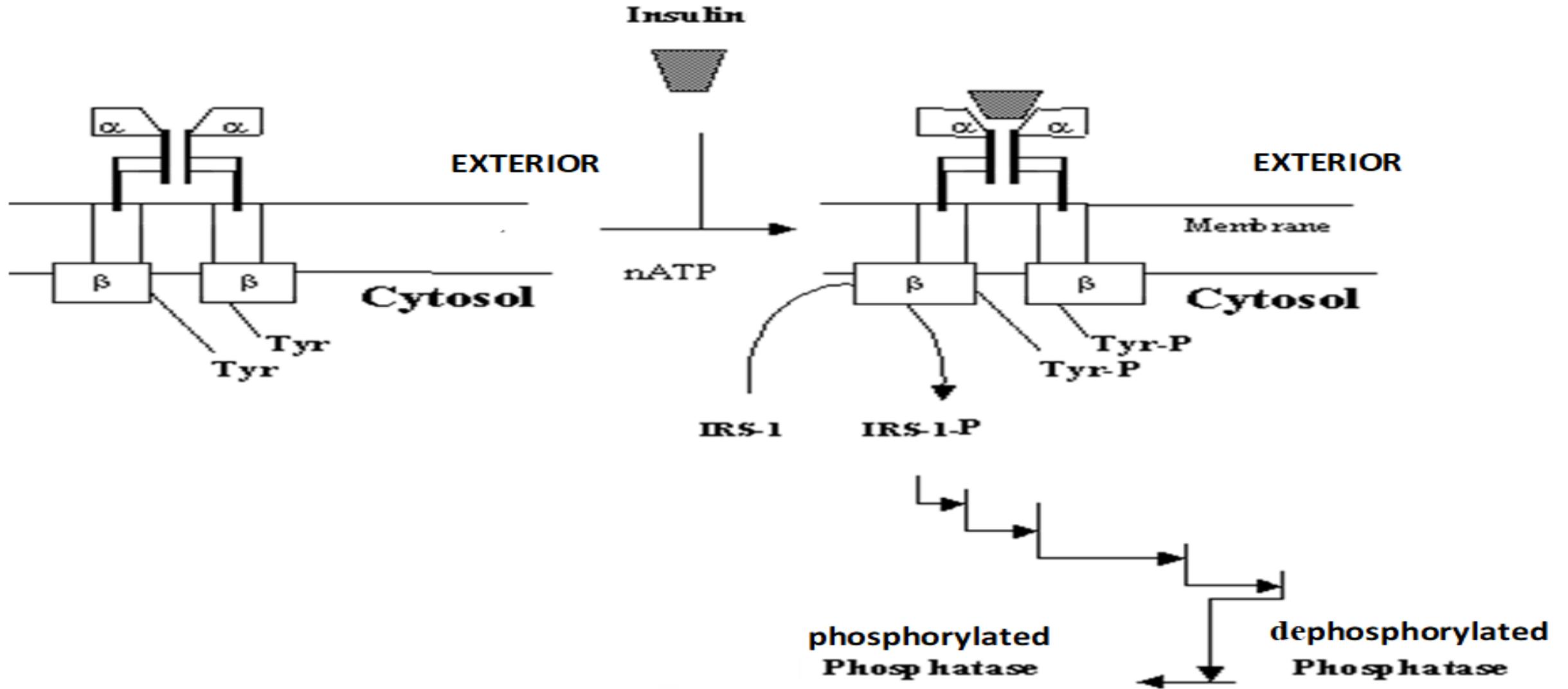


**Reaction 6: Glycogen branching**

# Glycogenesis



# Hormonal regulation of glycogenesis



Activation mechanism of protein phosphatase by insulin

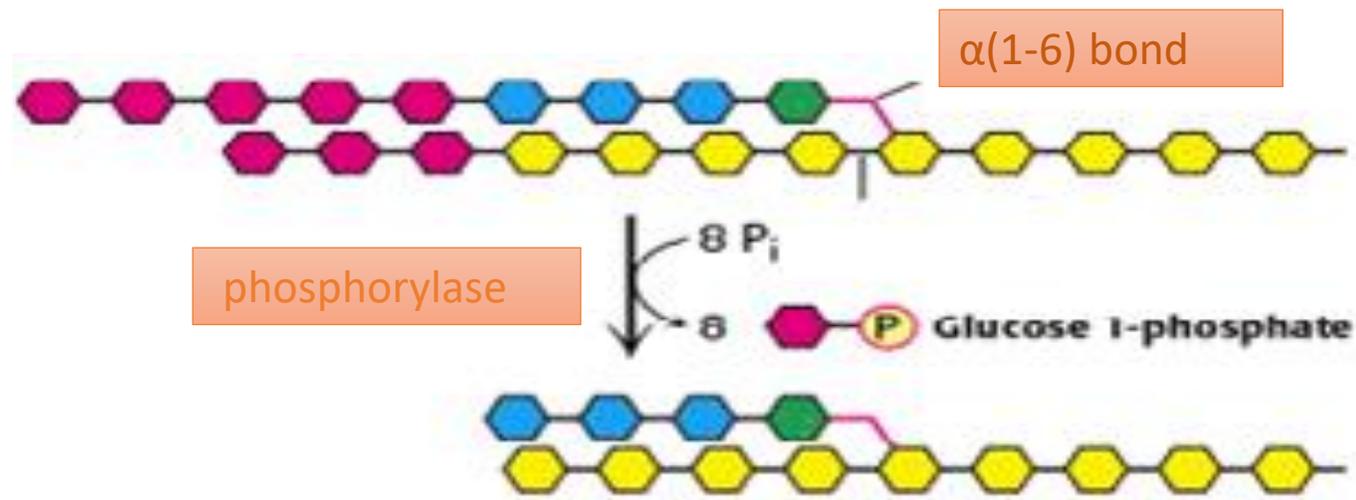
# Glycogenolysis reactions



# Glycogenolysis

## Reaction 1: Glycogen phosphorylase

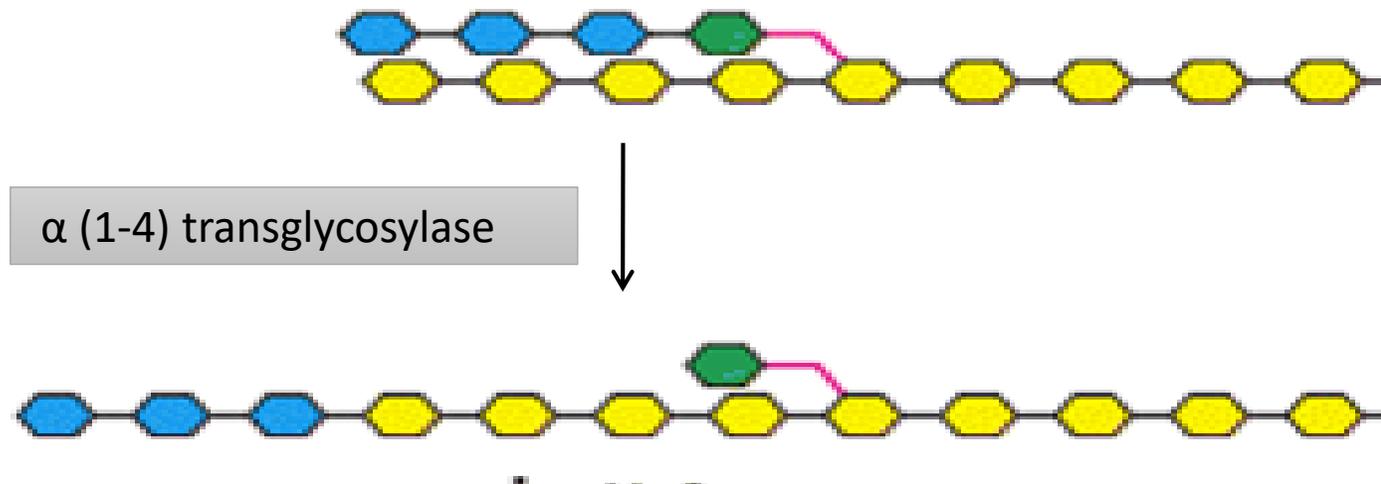
- Cleavage of the  $\alpha$  (1-4) bond from the non-reducing end
- Attachment of a phosphate group provided by ATP to carbon 1 of the released glucose
- Phosphorolysis is repeated sequentially on glycogen until 4 glycosyl residues are encountered on each chain before the  $\alpha$  (1-6) bond



# Glycogenolysis

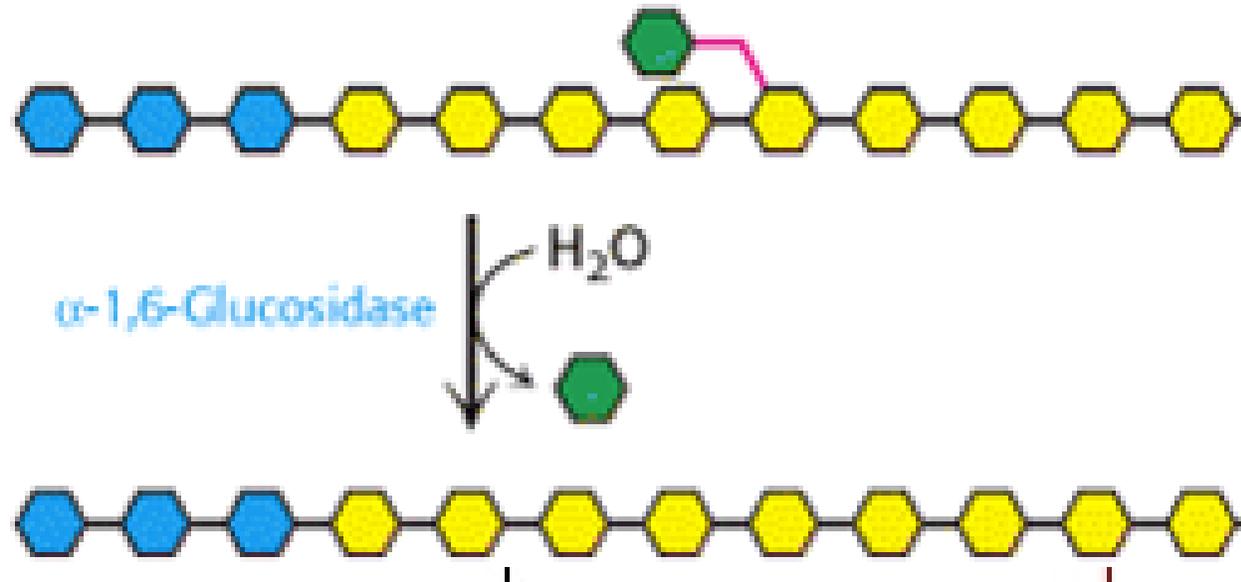
## Reaction2: glycosyl transferase effect by $\alpha$ (1-4) transglycosylase

- Glycosyltransferase acts on limiting dextrin by removing an oligoside made up of 3 glucose residues from each chain
- The three glucose residues are attached to another limit dextrin chain, thus allowing phosphorolysis to resume on this chain.
- After the action of this enzyme, a glucose molecule linked by the  $\alpha$  (1-6) bond remains in place of the side chain.



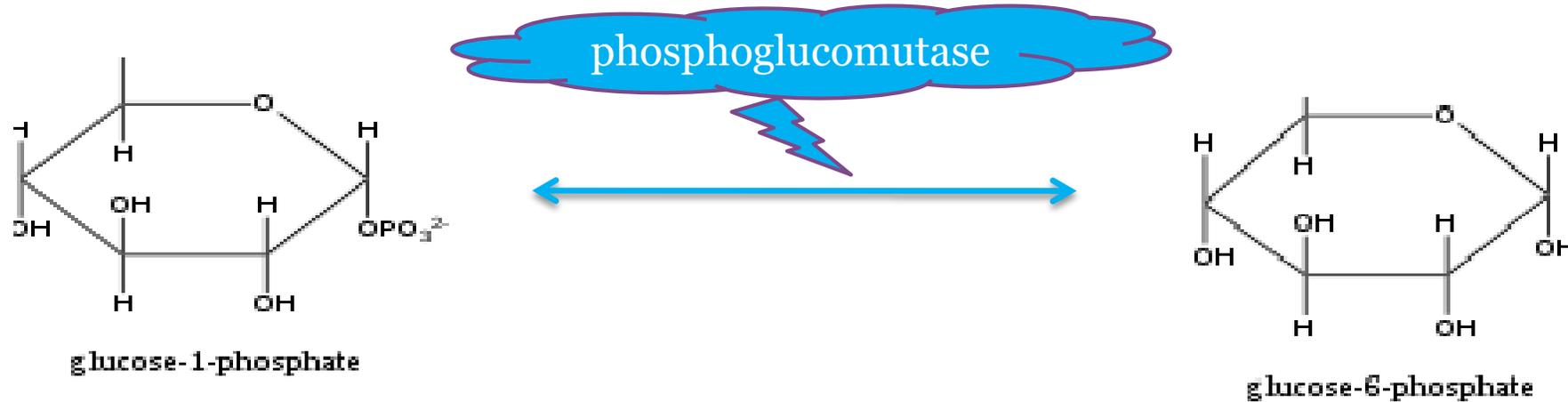
## Reaction 3: Debranching (ou $\alpha$ (1-6) glucosidase) effect

- A debranching enzyme hydrolyzes glucose residues linked by the  $\alpha$  (1-6) bond and releases glucose molecules

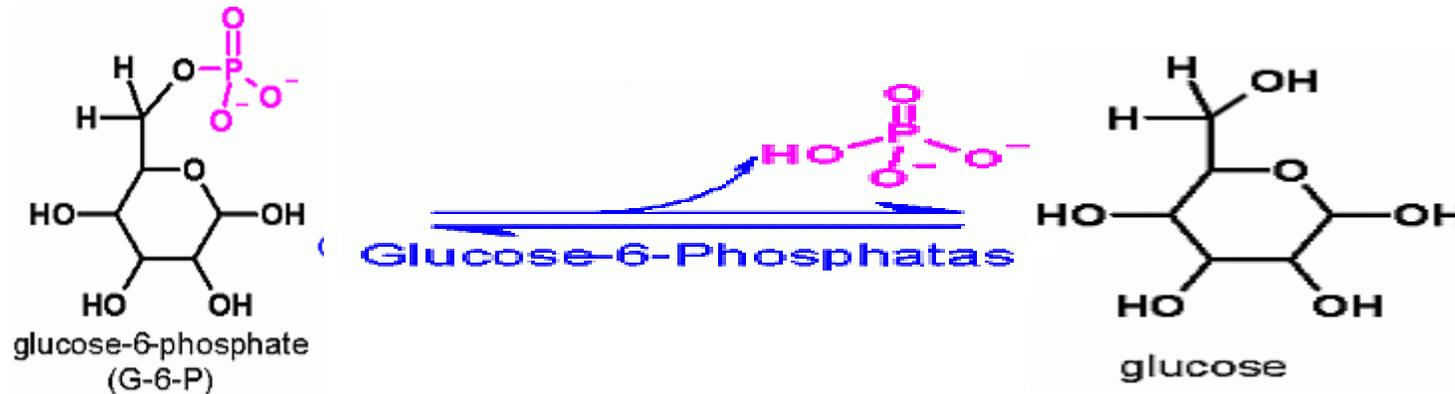


# Glycogenolysis

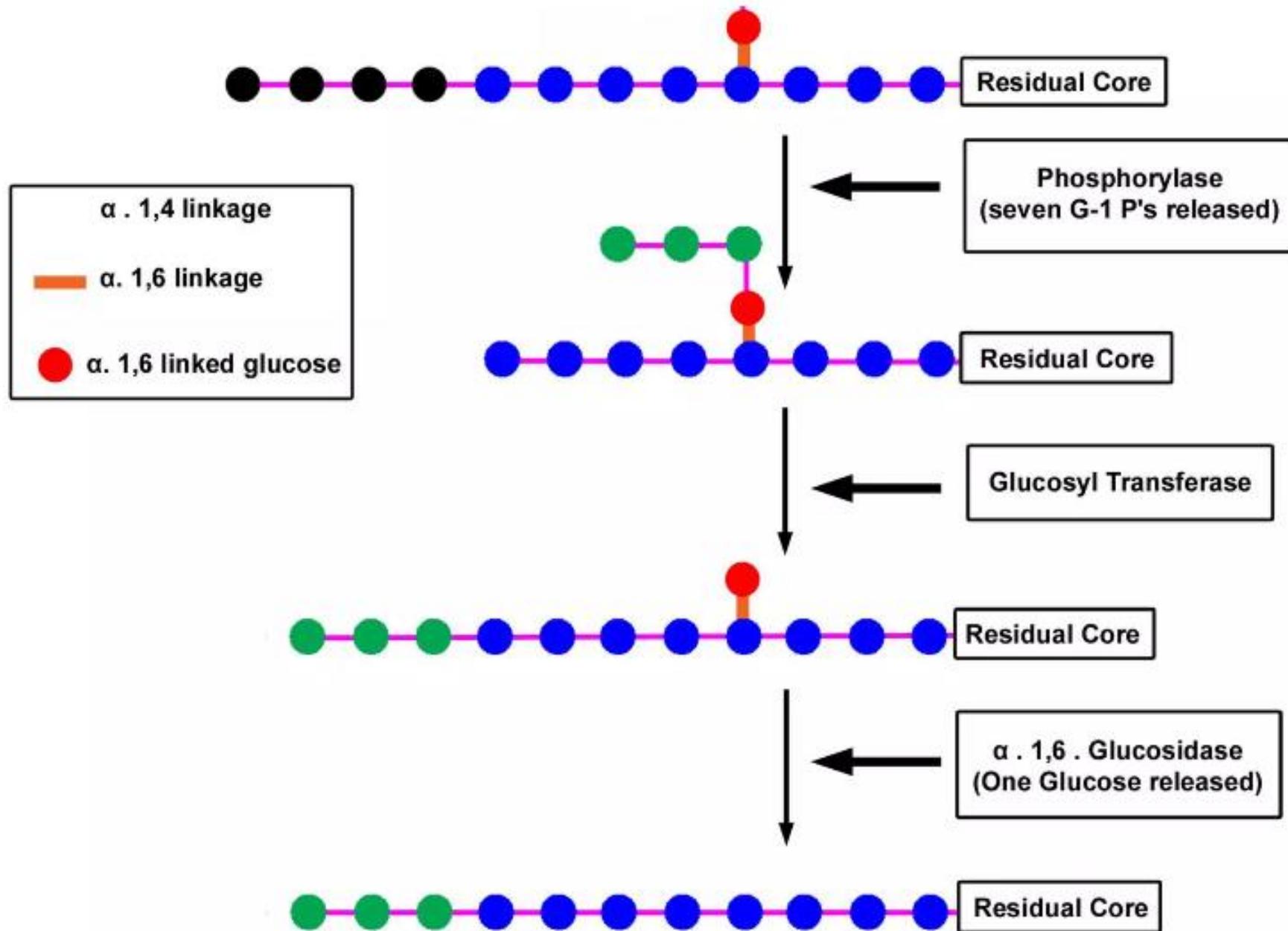
- G1P is converted to G6P by phosphoglucomutase. G6P can enter glycolysis in the liver and muscle



- Glucose 6-phosphatase allows the hydrolysis of G6P into glucose and the release of the latter into the blood



# Glycogenolysis



# Hormonal regulation of glycogenolysis

The binding of each of the hormones to its specific membrane receptor leads to activation of membrane-bound adenylate cyclase (adenylate

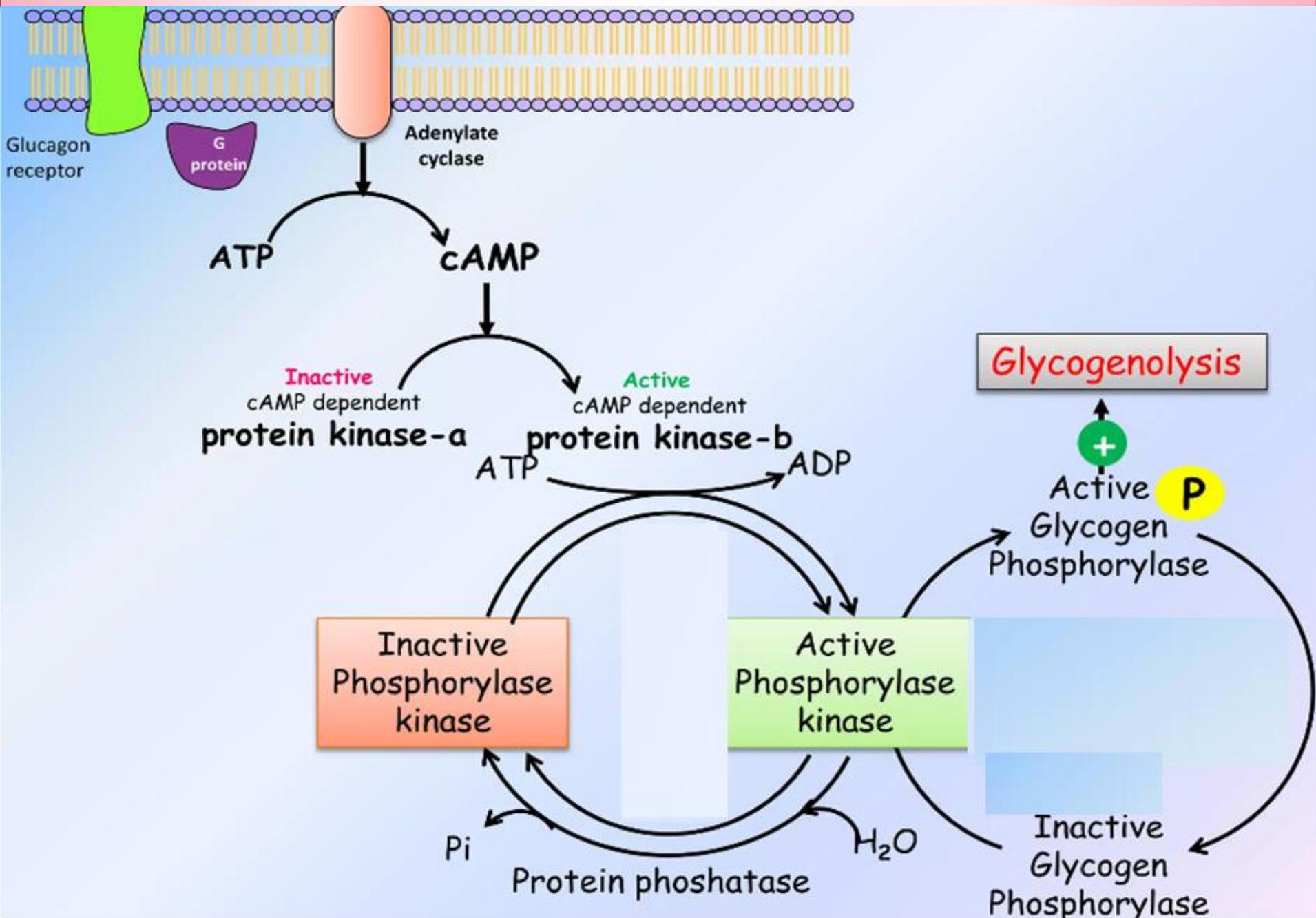
Activated adenylate cyclase catalyzes, through hydrolysis of ATP, the formation of cAMP from ATP. cAMP

kinase dependent combination of regulatory subunits and Kinase

Protein kinase A (active) phosphorylates glycogen phosphorylase which is in phosphorylated form.

Finally, this last one phosphorylates glycogen phosphorylase, changing it from the b form to the a form, which catalyzes the phosphorolysis of glycogen.

# Hormonal regulation of degradation of glycogen



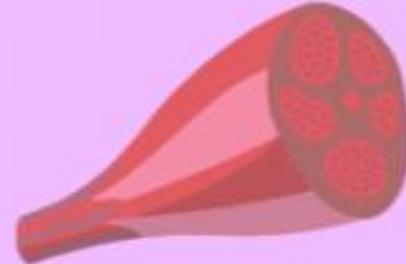
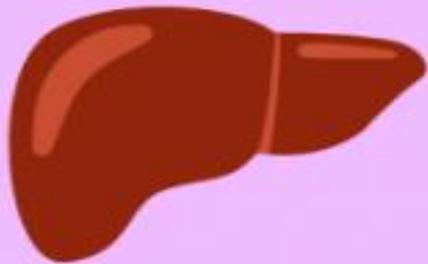
# Gluconeogenesis

Gluconeogenesis is the formation of glucose from non-carbohydrate carbon sources.

Fasting, Low Carb Diet, Intense Exercise, Starvation

The body uses carbon from fats and proteins to make sugar.

Amino Acids, Lactate, Glycerol



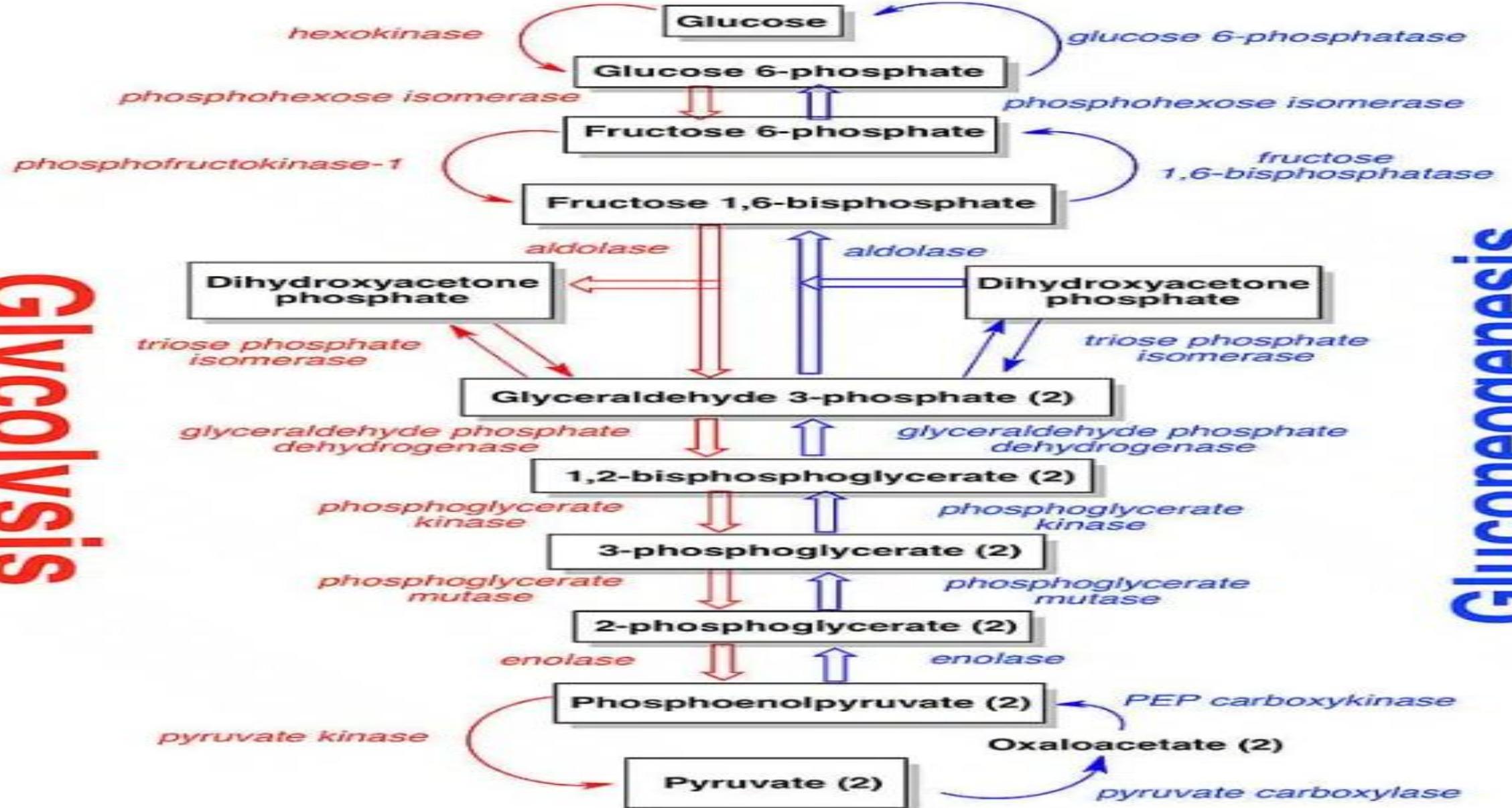
In humans, the process mainly occurs in the liver, but also in the kidney, intestine, muscle and brain.

**Glucose**

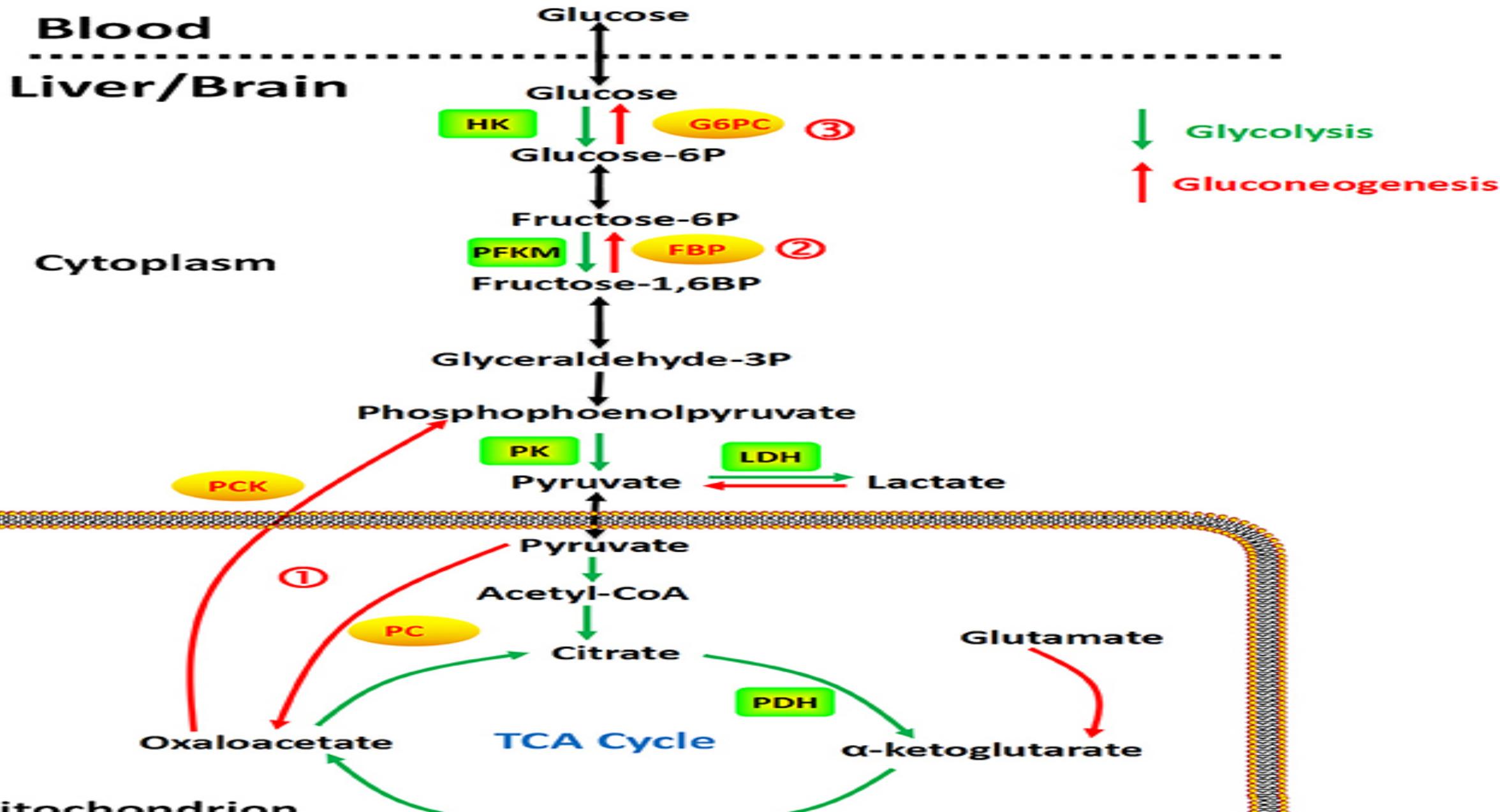
# Gluconeogenesis

Glycolysis

Gluconeogenesis

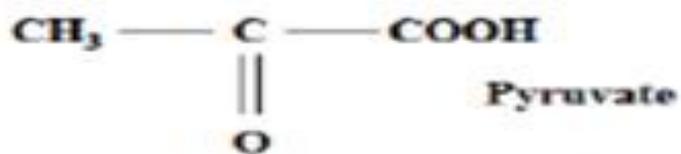


# Gluconeogenesis

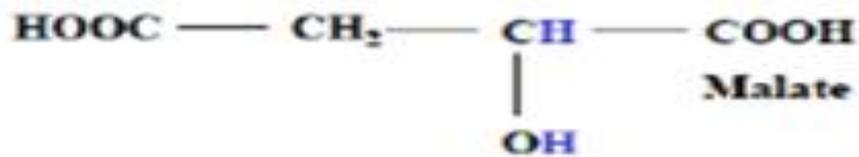
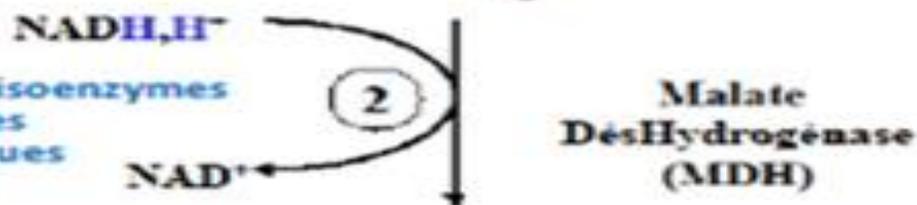


## Formation du phosphoénolpyruvate

Transport du pyruvate  
du cytoplasme  
vers mitochondrie

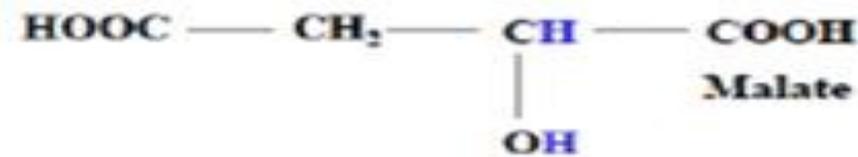
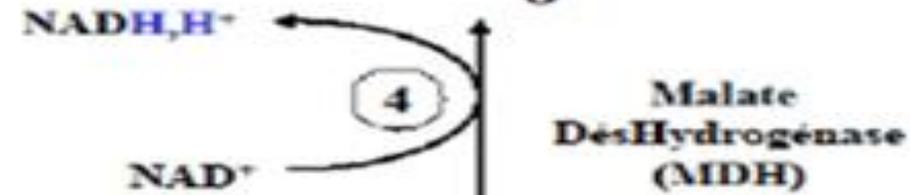
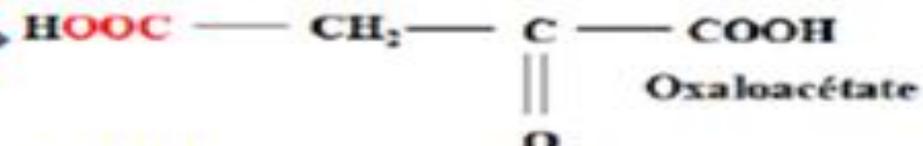
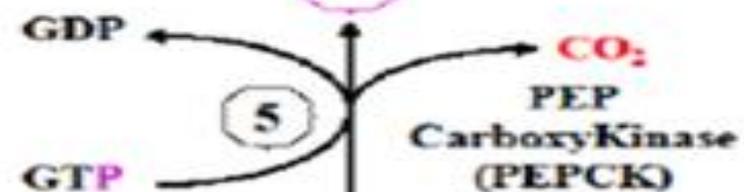
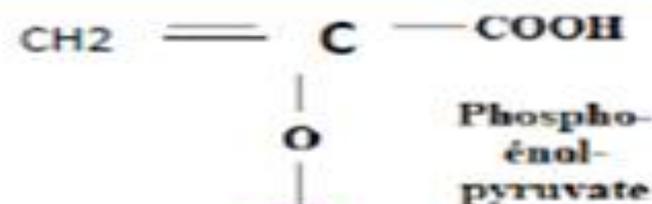


L'OAA ne peut traverser  
membrane mitochondriale



utilisation des isoenzymes  
mitochondriales  
et cytoplasmiques  
de la MDH

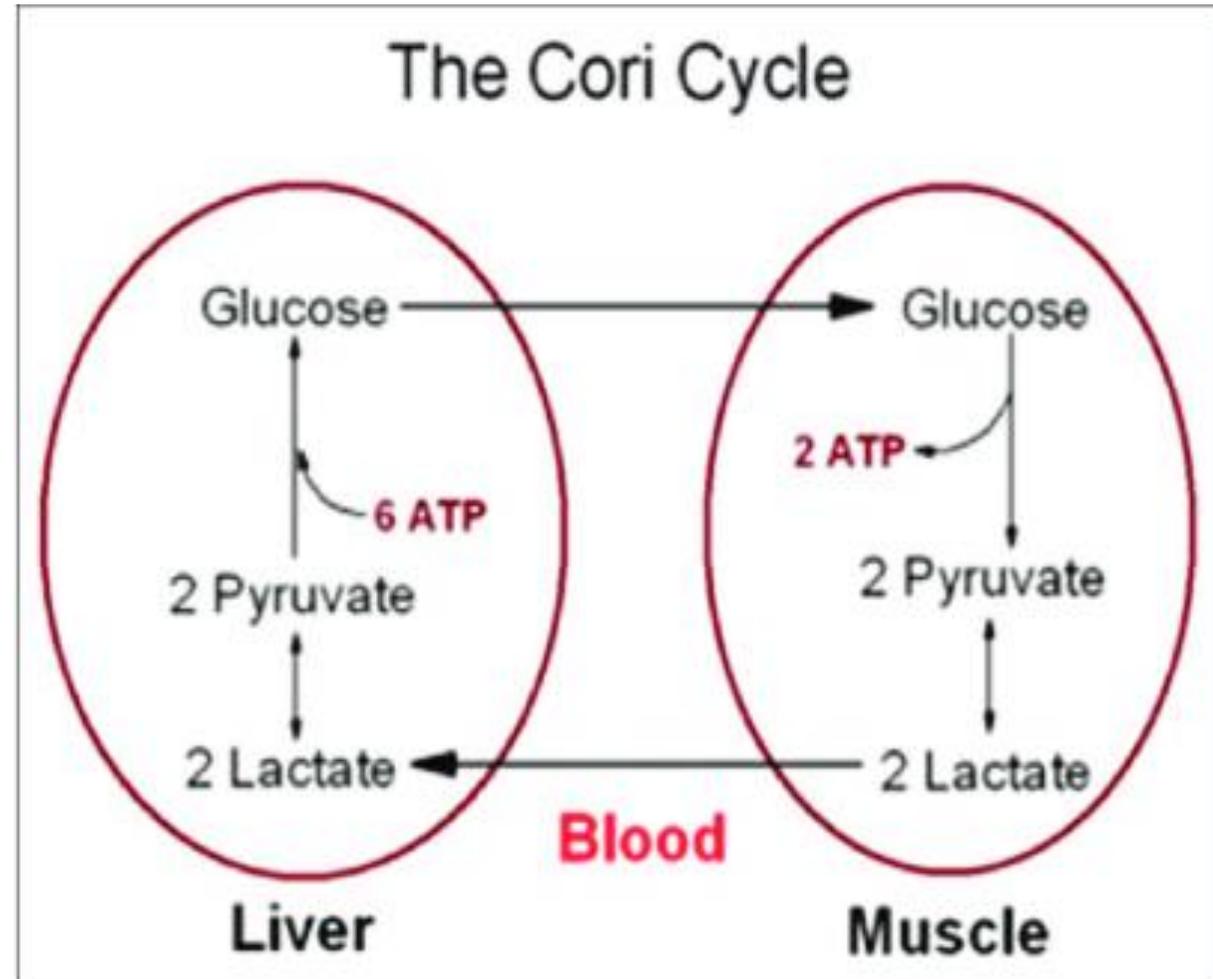
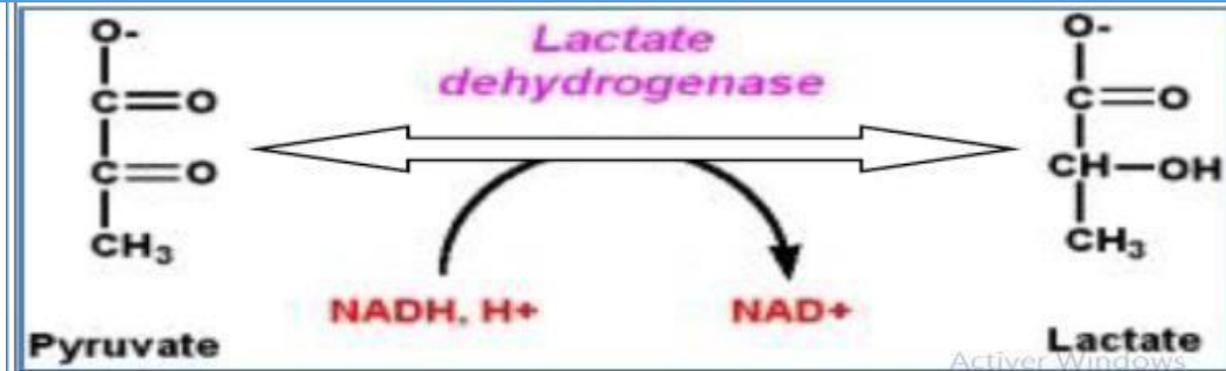
CO<sub>2</sub> Biotine



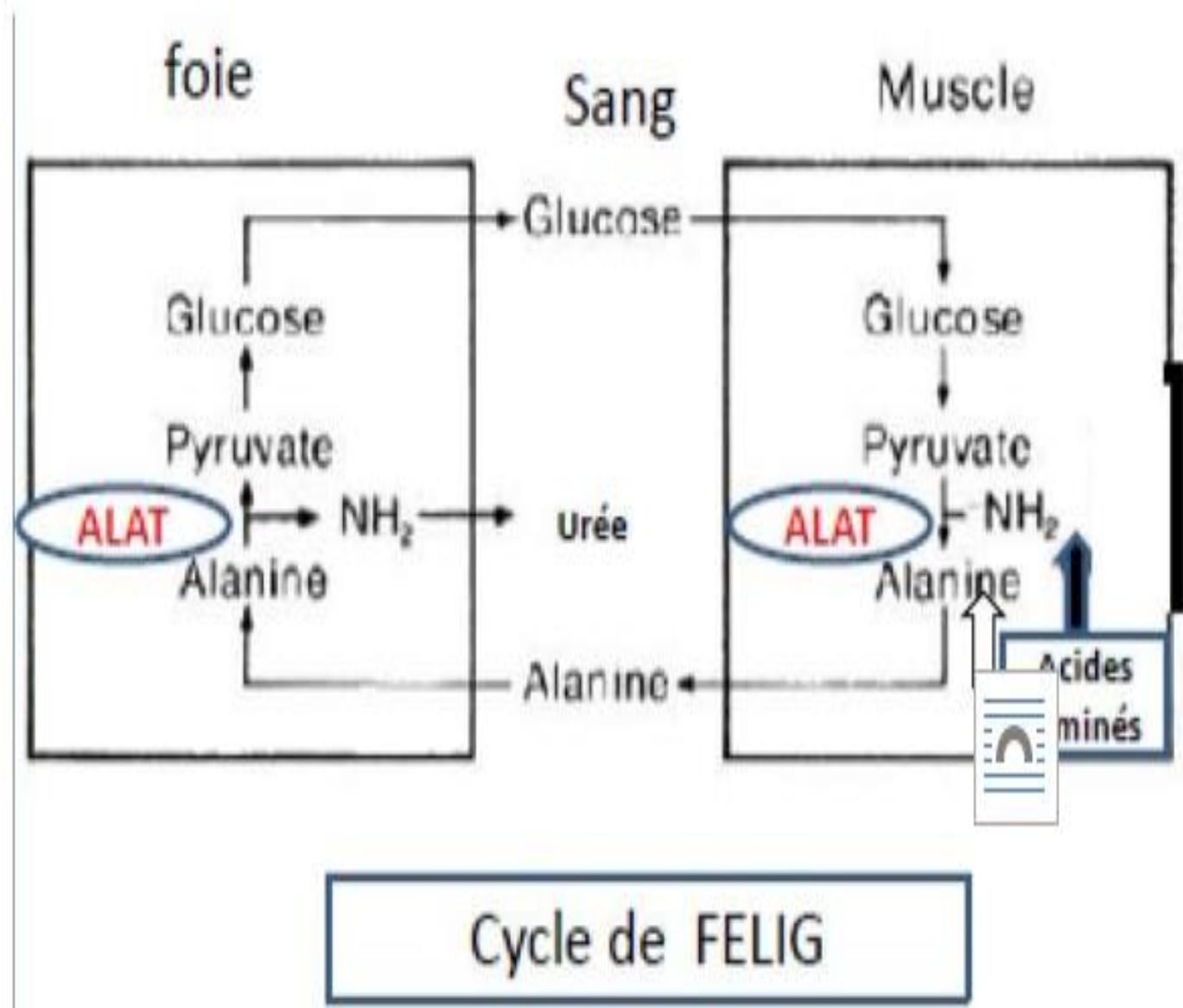
Navette  
malate

③

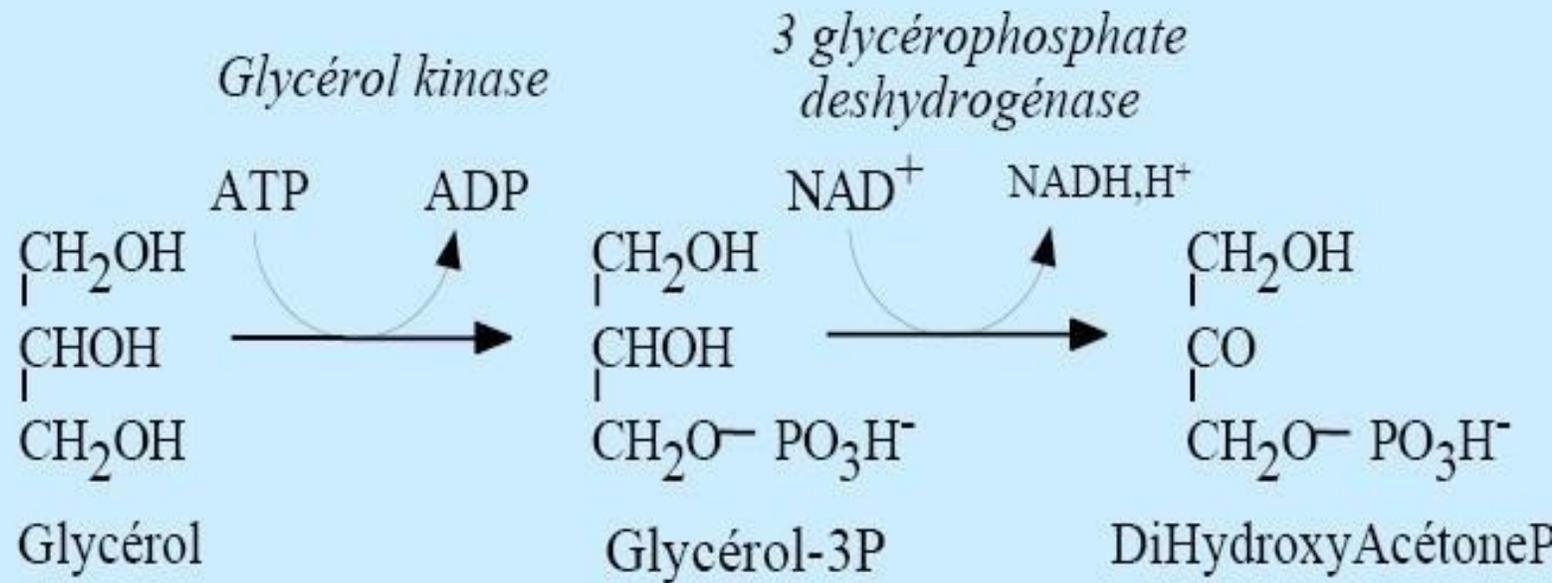
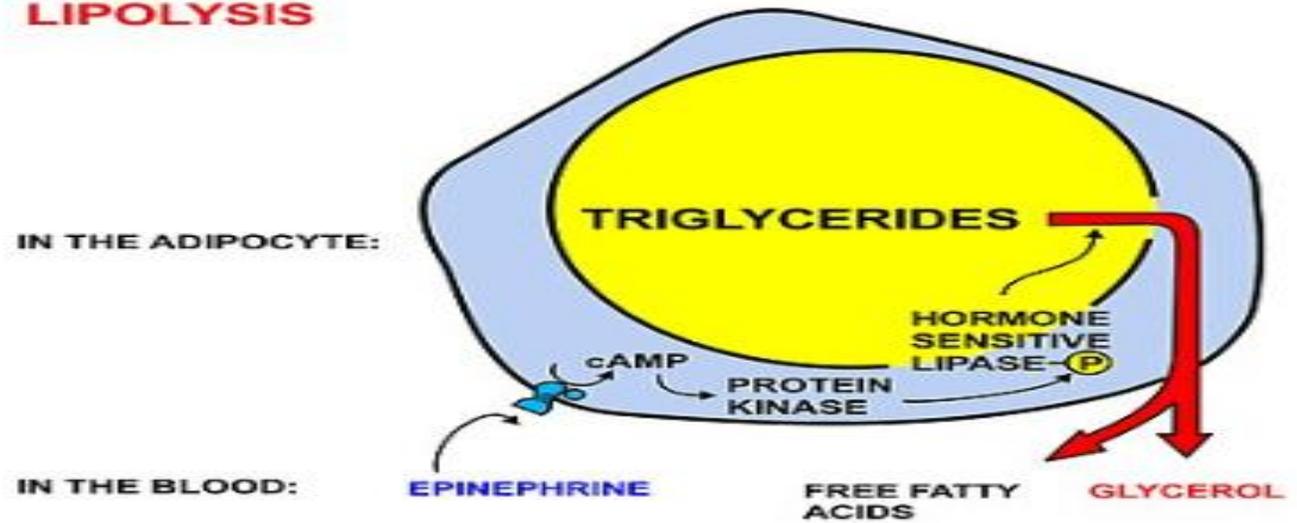
The Cori cycle is a crucial metabolic pathway that allows for the recycling of lactate produced in muscles and red blood cells, converting it back into glucose in the liver. This process is particularly important during intense exercise or when the oxygen supply is insufficient (anaerobic metabolism).



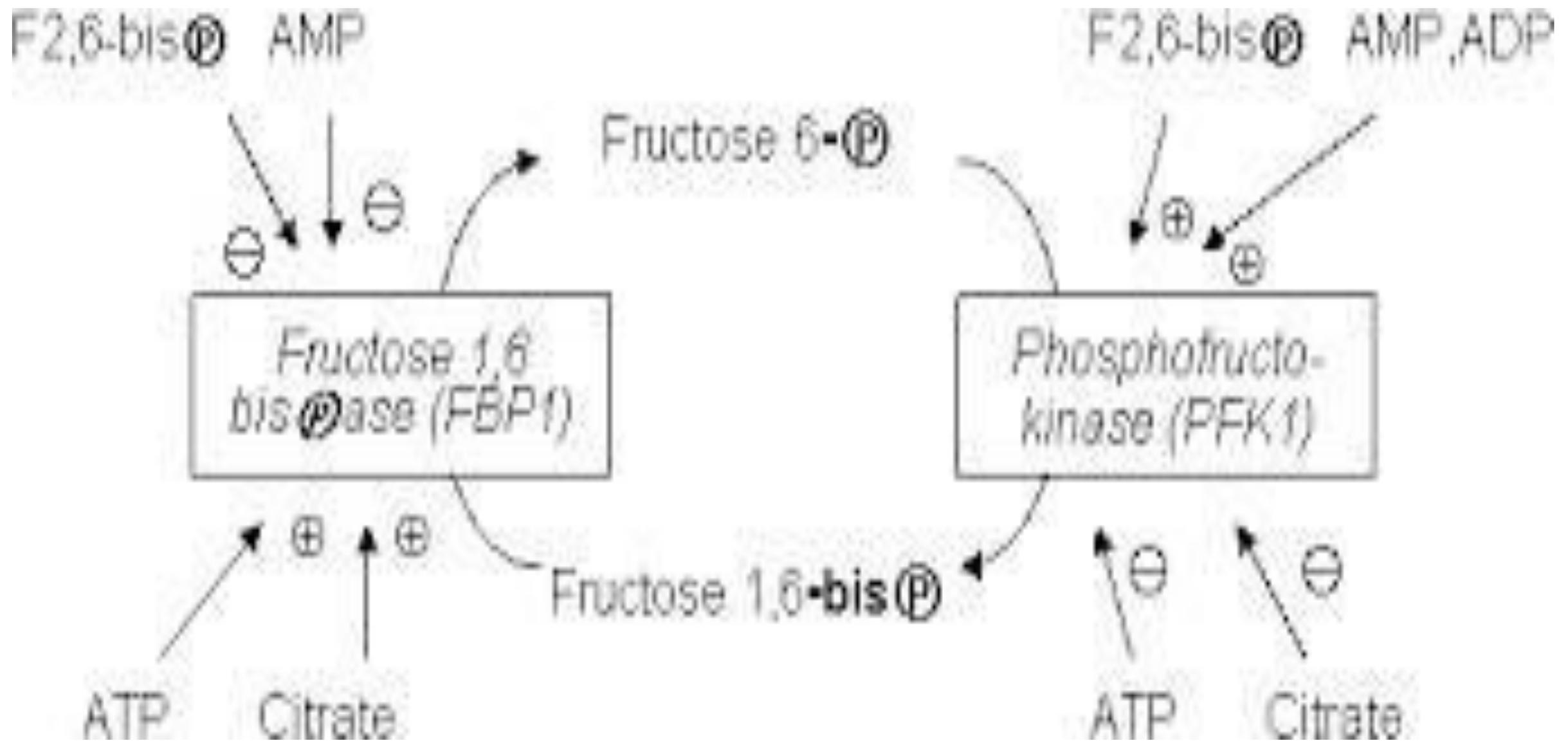
The **Felig cycle**, also known as the **glucose-alanine cycle**, is a metabolic pathway that ensures the transfer of nitrogen and carbon between skeletal muscles (and other peripheral tissues) and the liver, primarily during fasting or prolonged exercise.



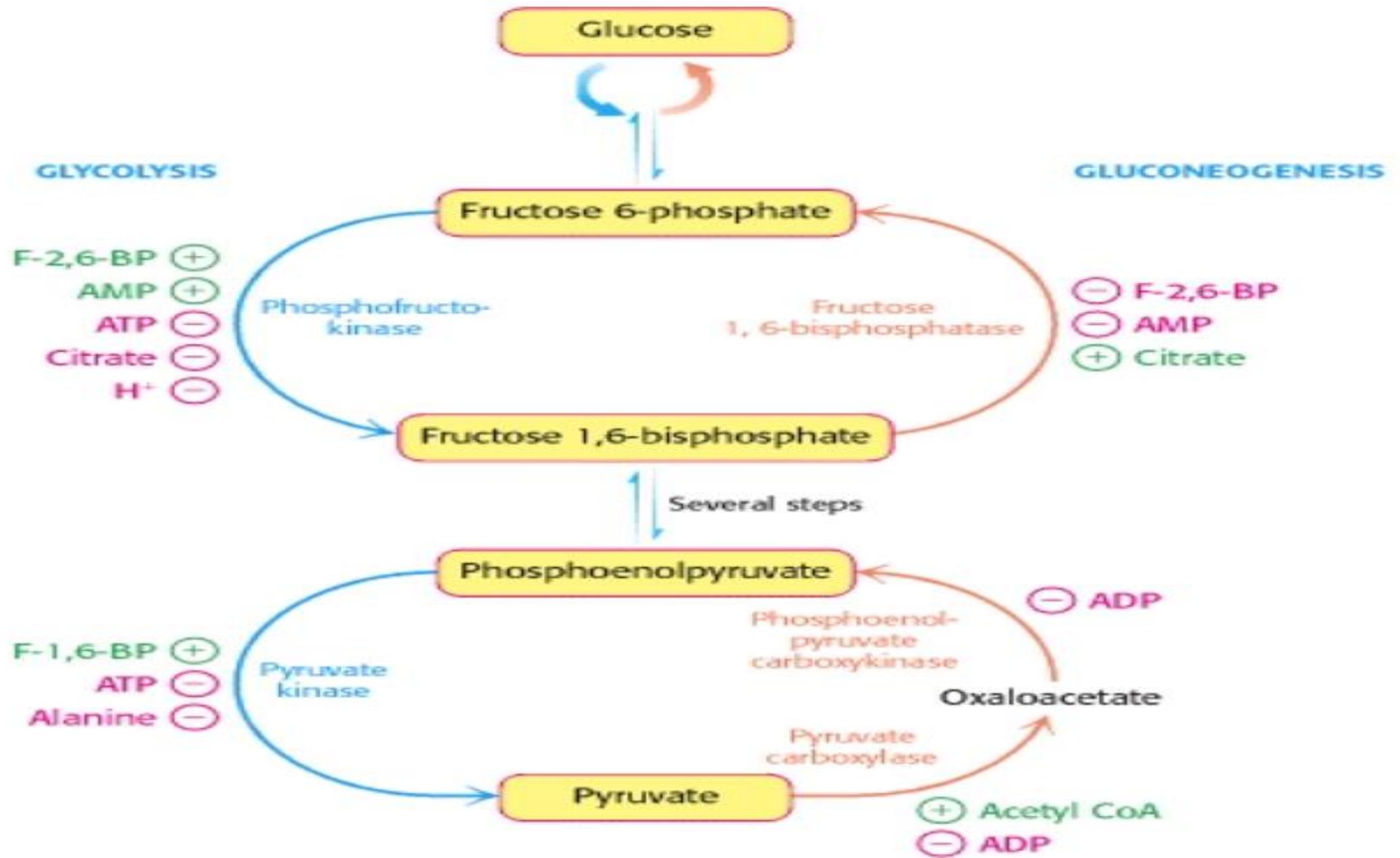
## LIPOLYSIS



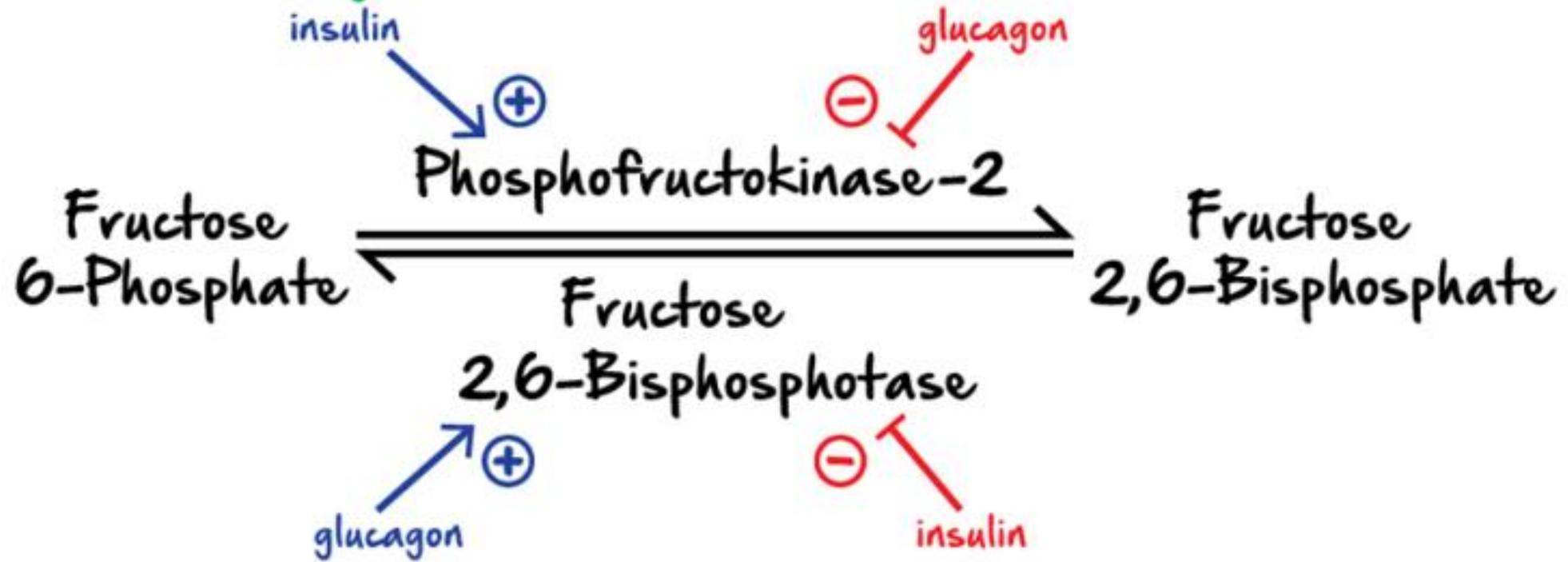
# Allosteric regulation of Gluconeogenesis



# Allosteric regulation of gluconeogenesis



## Hormonal Regulation



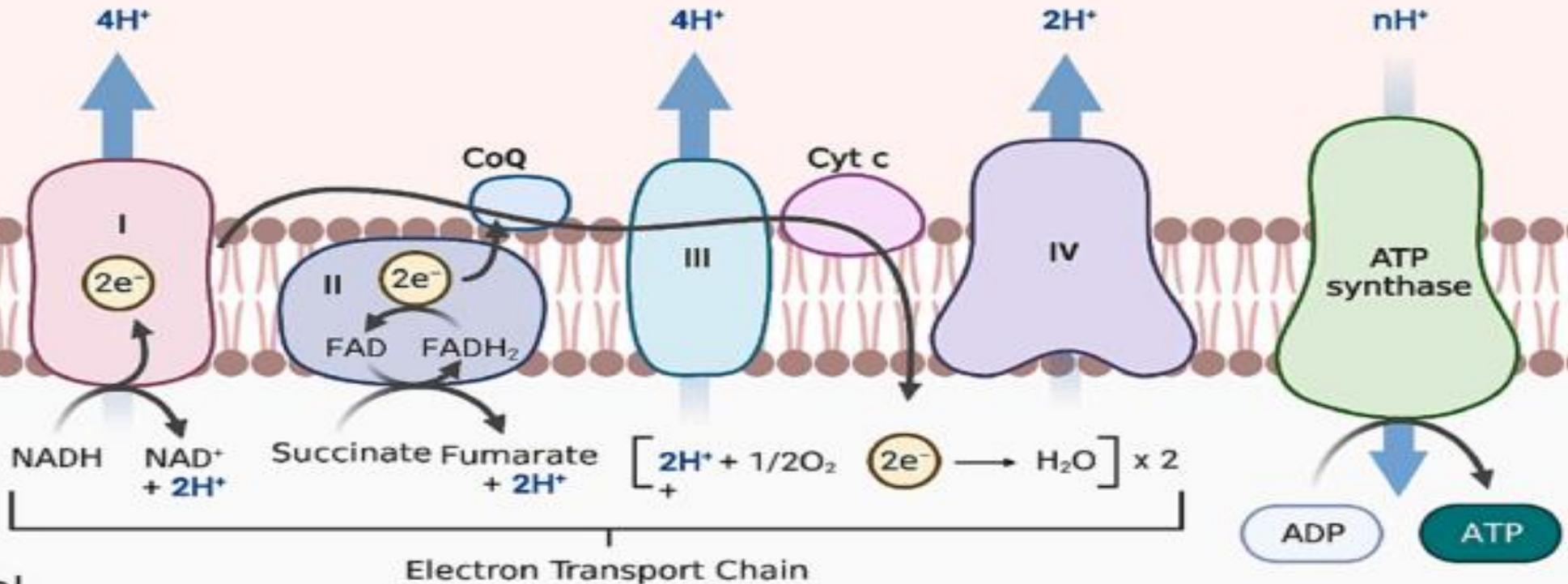
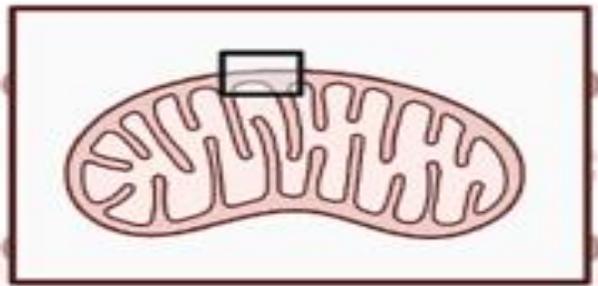
Cytoplasm

Outer membrane

Intermembrane space

Inner membrane

Mitochondrial matrix



Electron Transport Chain

Complex

I

II

III

IV

V