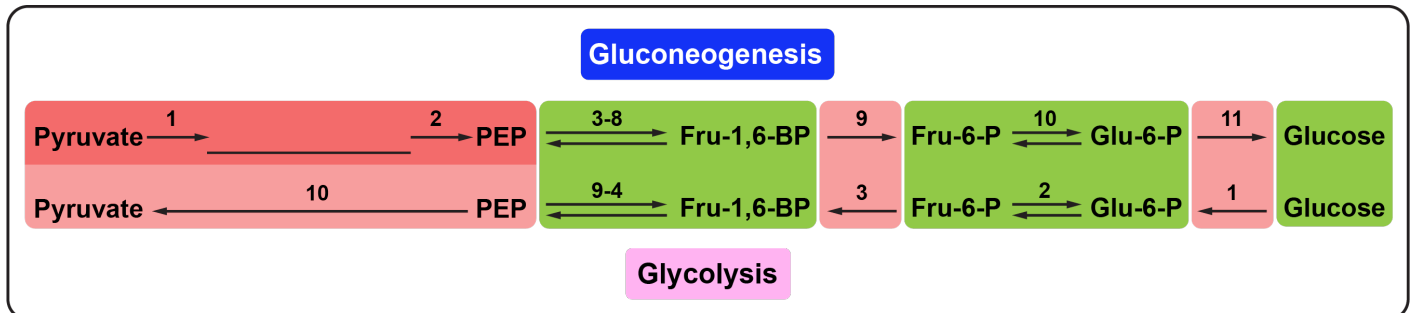


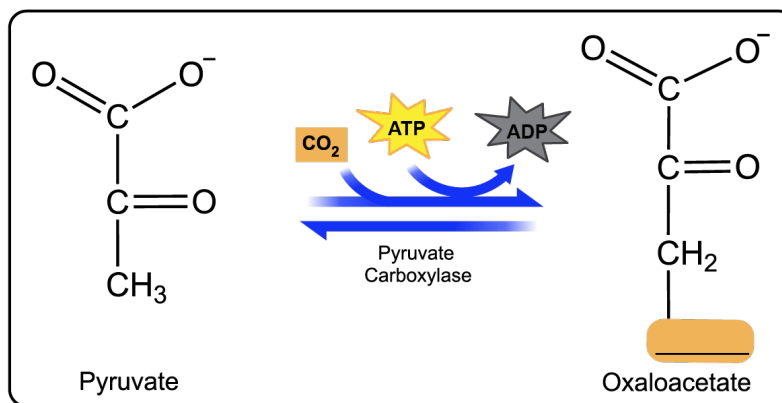
## CONCEPT: GLUCONEOGENESIS

- Gluconeogenesis is a sequence of \_\_\_\_ biochemical reactions with 2 pyruvates as the starting metabolites.
  - Reactions \_\_\_\_, & \_\_\_\_ are different from glycolysis, the rest are the **same**.



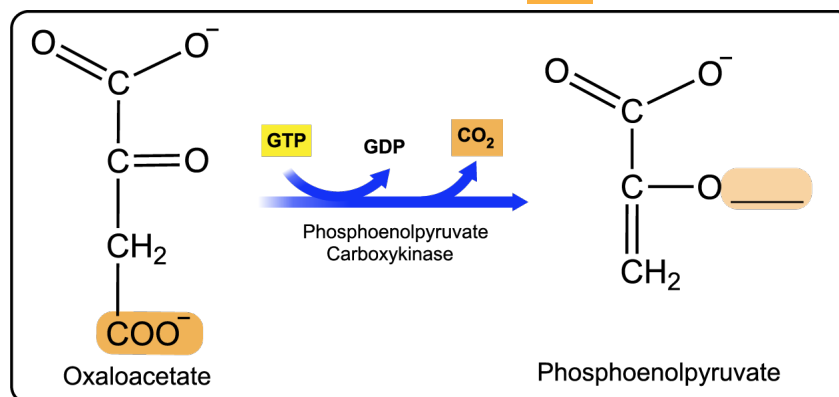
- 1 Oxaloacetate Formation x 2:**  $\text{CO}_2$  is added to pyruvate by pyruvate \_\_\_\_\_ lase to produce oxaloacetate.

- **ATP** is converted to \_\_\_\_\_.
- **Note:** \_\_\_\_\_ acids can enter as either pyruvate or oxaloacetate.



- 2 Decarboxylation & Phosphorylation x 2:** PEP carboxy\_\_\_\_\_ removes  $\text{CO}_2$  and adds \_\_\_\_\_ group.

- **GTP** is converted to GDP.
- 1 C atom is lost as  **$\text{CO}_2$** .



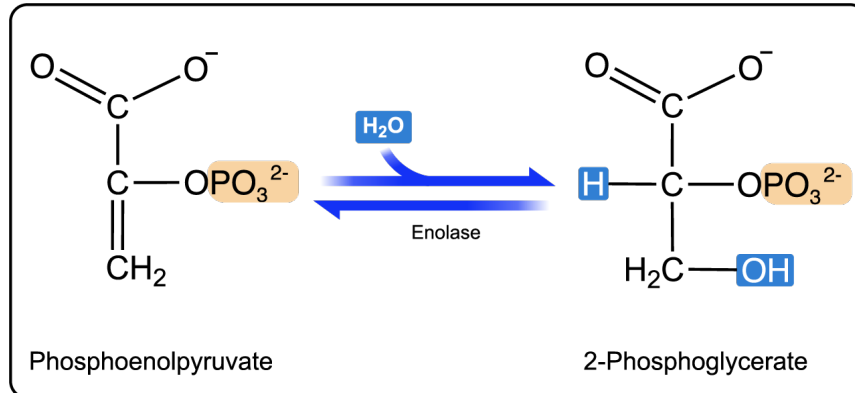
**EXAMPLE:** How many total carbon atoms are lost in the first 2 reactions of gluconeogenesis?

- a) 0                      b) 1                      c) 2                      d) 4

## CONCEPT: GLUCONEOGENESIS

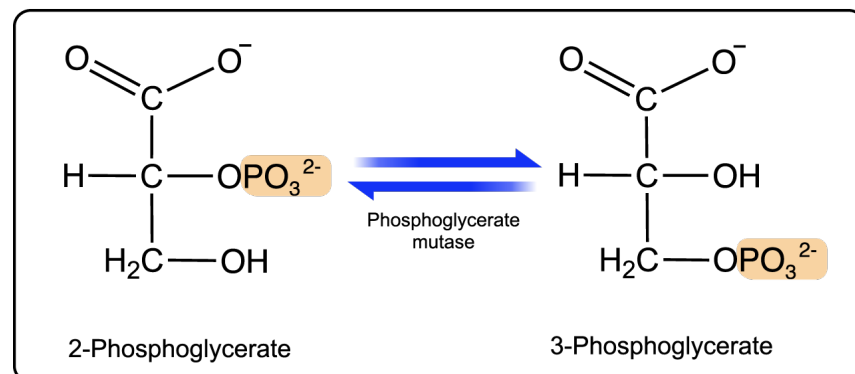
**3 Hydration x 2:** PEP undergoes hydration to produce 2PG.

□ Catalyzed by enzyme \_\_\_\_\_.



**4 Isomerization x 2:** 2PG undergoes isomerization to yield 3PG.

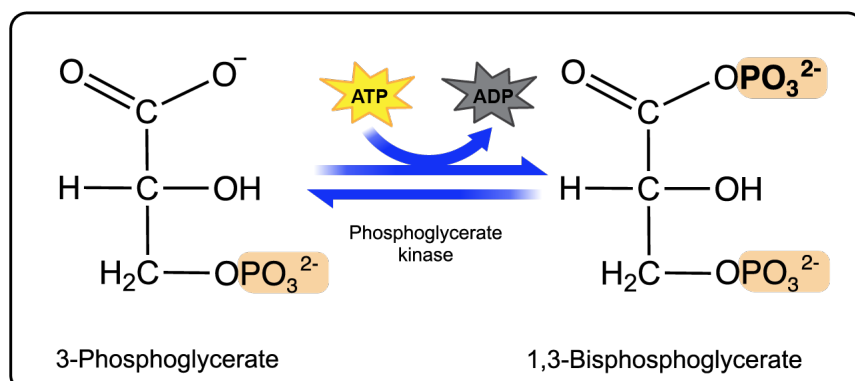
□ Catalyzed by enzyme phosphoglycerate \_\_\_\_\_.



**5 Phosphate Transfer x 2:** 3PG produces 1,3-bisphosphoglycerate by gaining a P<sub>i</sub> group.

□ Catalyzed by phosphoglycerate \_\_\_\_\_.

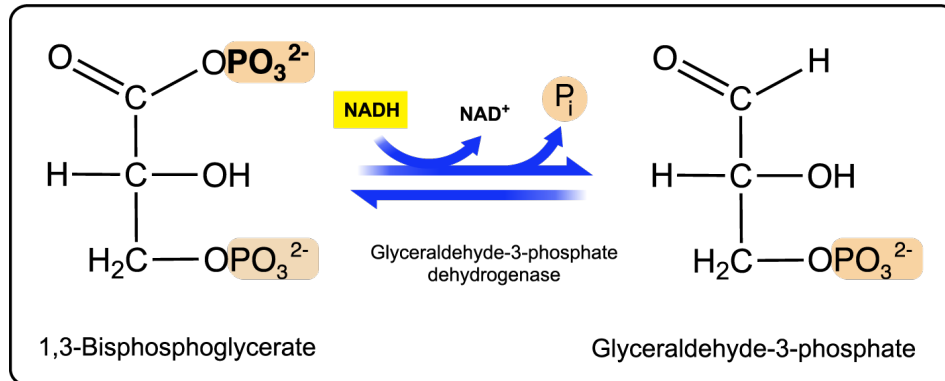
□ ATP is converted to ADP.



## CONCEPT: GLUCONEOGENESIS

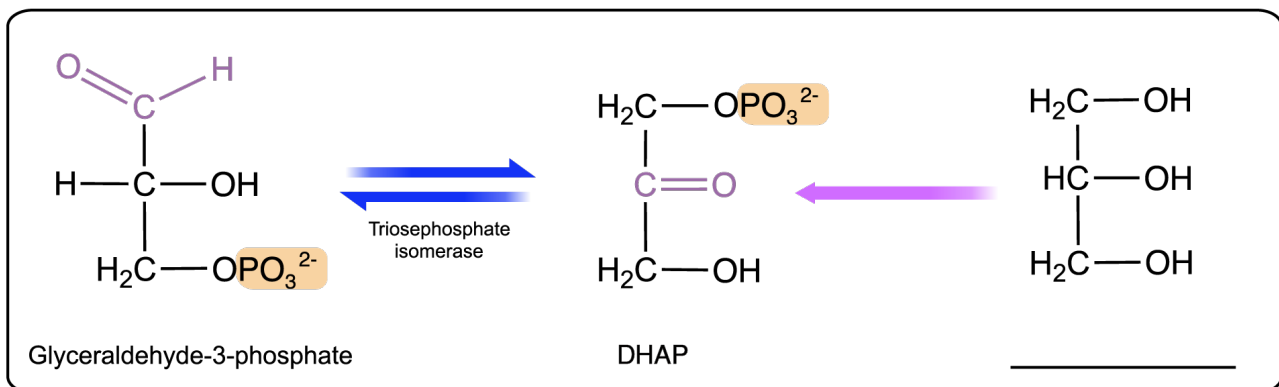
**6 Reduction x 2:** 1,3-bisphosphoglycerate undergoes reduction to produce G3P.

- ☐ Catalyzed by enzyme G3P \_\_\_\_\_. ☐ **NADH** is oxidized to  $\text{NAD}^+$ .

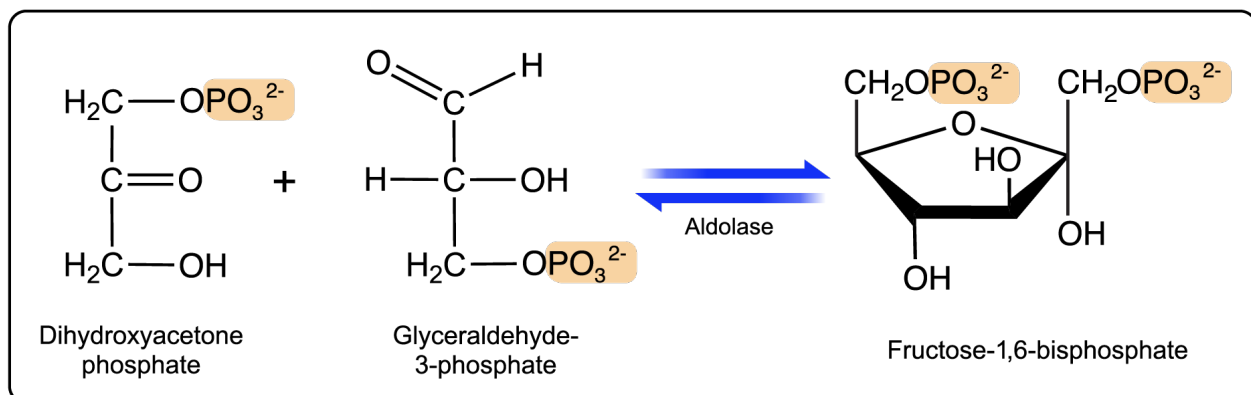


**7 Isomerization:** One G3P molecules is isomerized to dihydroxyacetone phosphate (DHAP).

- ☐ Catalyzed by enzyme triosephosphate \_\_\_\_\_.  
☐ **Note:** Glycerol enters gluconeogenesis as \_\_\_\_\_.



**8 Linkage:** Enzyme \_\_\_\_\_ combines 2 triose phosphates into fructose-1,6-bisphosphate.

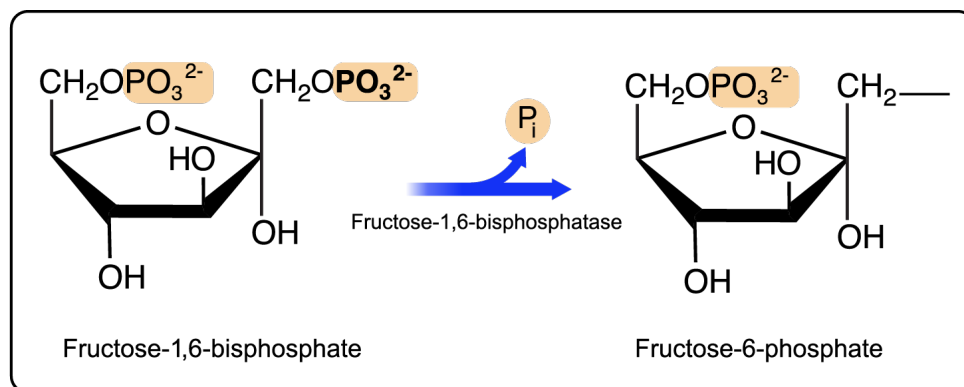


## CONCEPT: GLUCONEOGENESIS

**EXAMPLE:** Which enzymes so far are involved in glycolysis but not gluconeogenesis?

- a) pyruvate carboxylase
- b) phosphoglycerate kinase
- c) pyruvate kinase
- d) glyceraldehyde-3-phosphate dehydrogenase

**9 Dephosphorylation:** Enzyme fructose-1,6-bisphosphatase removes a  $P_i$  group to form fructose-6-phosphate.

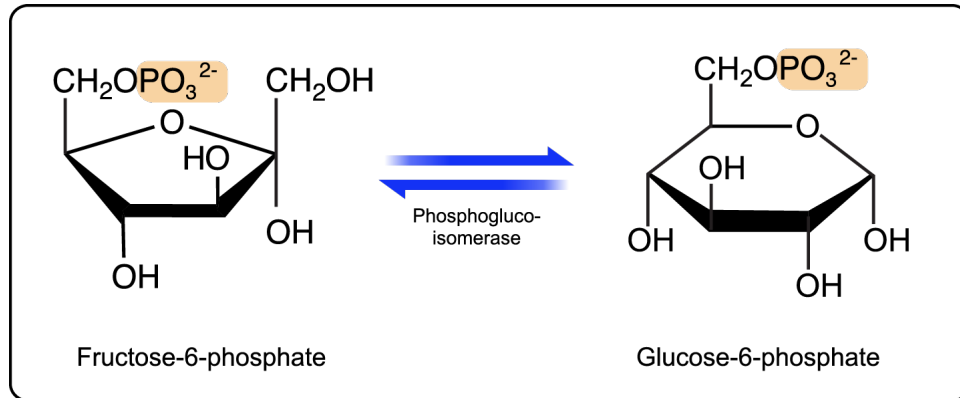


**EXAMPLE:** Why is it not possible to dephosphorylate fructose-1,6-bisphosphate with phosphofructokinase enzyme?

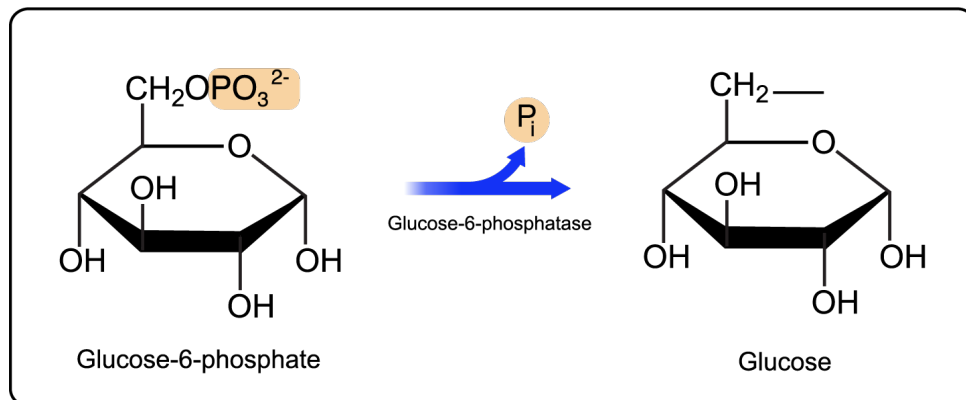
- a) Phosphofructokinase does not function properly in the cytoplasm of the cell.
- b) Phosphofructokinase catalyzes transfer of a  $P_i$  group from one molecule to another, not removal of a  $P_i$  group.
- c) Phosphofructokinase lacks enough energy to catalyze dephosphorylation.
- d) It is possible.

## CONCEPT: GLUCONEOGENESIS

**10 Isomerization:** Enzyme phosphogluco\_\_\_\_\_ isomerizes fructose-6-phosphate into glucose-6-phosphate.



**11 Dephosphorylation:** Enzyme glucose-6-phosphatase removes a  $\text{P}_i$  group forming \_\_\_\_\_.



**EXAMPLE:** Step 9 and 11 of Gluconeogenesis:

- a) Uses ADP to dephosphorylate and produces ATP.
- b) Produces ADP as a result of dephosphorylation.
- c) Requires GTP to dephosphorylate.
- d) Does not require energy to dephosphorylate.

## CONCEPT: GLUCONEOGENESIS

### Remembering Gluconeogenesis

Remembering Gluconeogenesis			
<b>MEMORY TOOL 1</b> (Reactions)	In the ____ <sup>st</sup> ____ minutes of accident	call ____	____
<b>MEMORY TOOL 2</b> (Metabolites)	____irates ____nly ____op	____ruit ____rom	____orgeous ____ardens
<b>MEMORY TOOL 3</b> (Enzymes)	____irates ____onsume _____-____	____ructose ____iz ____izz	____lucose ____izz

### Summary

- Gluconeogenesis reduces pyruvate to glucose via following reaction:



### MEMORY TOOL 4:

Reaction 1: **ATP**

\_\_\_\_ll \_\_\_\_he \_\_\_\_irates

Reaction 2: **GTP**

\_\_\_\_ot \_\_\_\_o \_\_\_\_arty

Reaction 5: **ATP**

\_\_\_\_ 5 \_\_\_\_m

**EXAMPLE:** Reactions 1 & 2 consume \_\_\_\_ & \_\_\_\_ and are catalyzed by \_\_\_\_ & \_\_\_\_.

- ATP & GTP, Pyruvate oxaloacetase & Oxaloacetate carboxykinase.
- ATP & GTP, Pyruvate carboxylase & Phosphoenolpyruvate carboxykinase.
- GTP & ATP, Pyruvate carboxykinase & Phosphoenolpyruvate carboxylase.
- GTP & ATP, Phosphoenolpyruvate carboxykinase & Phosphoglycerate kinase.

**CONCEPT: GLUCONEOGENESIS**

**PRACTICE:** The following metabolites are present both in glycolysis and gluconeogenesis, except:

- a) oxaloacetate
- b) fructose-1,6-bisphosphate
- c) glyceraldehyde-3-phosphate
- d) dihydroxyacetone phosphate

**PRACTICE:** Which molecule is added to pyruvate and then removed in the next reaction?

- a)  $H^+$
- b)  $H_2O$
- c)  $PO_3^{2-}$
- d)  $CO_2$

**PRACTICE:** Enzyme responsible for decarboxylation and phosphate transfer in the same gluconeogenic reaction is:

- a) Tyrosine kinase
- b) Oxaloacetate carboxykinase
- c) Pyruvate kinase
- d) Phosphoenolpyruvate carboxykinase
- e) Phosphofructokinase

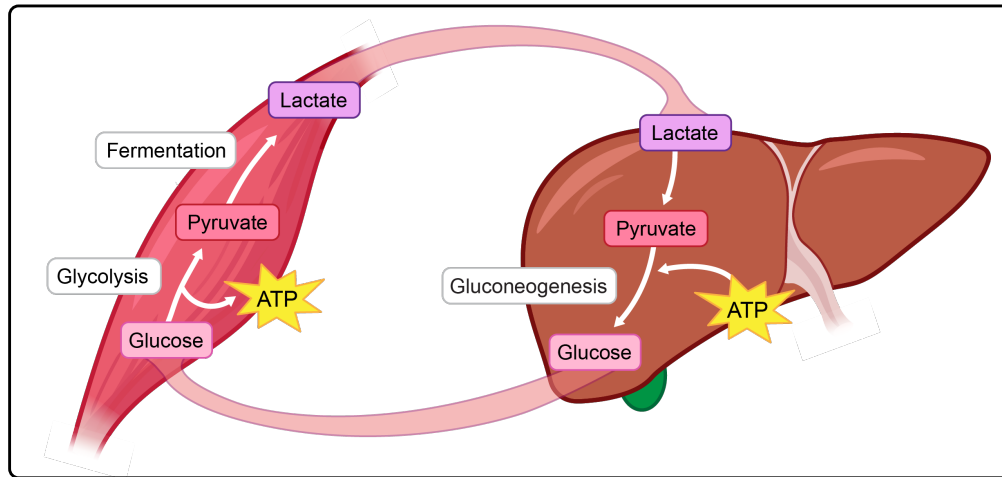
**PRACTICE:** Fructose-1,6-bisphosphate is dephosphorylated by \_\_\_\_\_ to fructose-6-phosphate.

- a) Glucose-6-phosphatase
- b) Fructose-1,6-bisphosphatase
- c) Phosphofructokinase
- d) Fructose-6-phosphatase

## CONCEPT: GLUCONEOGENESIS

### Cori Cycle

- Cyclic metabolic pathway that transports \_\_\_\_\_ from muscle cells to liver cells and converts it to \_\_\_\_\_.
- **Recall:** Lactate is produced by muscle cells during \_\_\_\_ aerobic conditions.
  - It is then transported by bloodstream to liver cells and converted to glucose through \_\_\_\_\_ neogenesis.



**EXAMPLE:** The primary purpose for Cori cycle is to:

- Produce lactate through fermentation process.
- To provide a metabolite for gluconeogenesis.
- Generate glucose for muscle cells to use as energy source.
- Provide liver with much needed energy.

**PRACTICE:** Which of the following statement(s) correctly describes the Cori cycle?

- Lactate is converted back to pyruvate in the muscles.
- Conversion of lactate to glucose in the liver generates ATP.
- Glucose from muscle cells is transported to liver through bloodstream.
- Lactate from the muscle cells is regenerated into glucose in the liver.
- Both a & b.
- a, c & d