## **Glycolysis Reactions**

- Glycolysis is a sequence of \_\_\_\_ biochemical reactions.
  - □ Reactions \_\_\_ to \_\_\_ split 1 glucose molecule into 2 glyceraldehyde 3-phosphate (G3P) molecules.
  - □ Reactions \_\_\_ to \_\_\_ convert G3P to \_\_\_\_\_ and produce high-energy molecules (ATP and NADH).
- **1** Phosphorylation: C\_\_ OH of  $\alpha$ -glucose attacks the \_\_-phosphorous of ATP.

- **2 Isomerization:**  $\alpha$ -Glucose 6-phosphate isomerizes to  $\alpha$ -fructose 6-phosphate.
  - □ Isomerization takes place through an \_\_\_\_\_ intermediate.

- **3** Phosphorylation: α-Fructose 6-phosphate undergoes phosphorylation to yield  $\beta$ -fructose 1,6-bisphosphate.
  - 3a) Mutarotation occurs \_\_\_\_\_ phosphorylation. 3b □ Phosphorylation occurs at C\_ OH.

- 4 Bond Cleavage: Fructose 1,6-bisphophate undergoes a \_\_\_\_\_ reaction.
  - (4a) An iminium intermediate is formed first.

**4b)** C\_\_-C\_\_ bond of the intermediate is cleaved.

4c) and 4d) Tautomerization and \_\_\_\_\_\_ of the enamine produces dihydroxyacetone phosphate (DHAP).

- 5 Isomerization: Dihydroxyacetone phosphate (DHAP) is isomerized to glyceraldehyde-3-phosphate (G3P).
  - □ Occurs via \_\_\_\_\_ mechanism.

**EXAMPLE:** Identify the mechanism through which DHAP is converted into G3P.

- a) Phosphorylation
- b) Enediol mechanism
- c) E1cB elimination
- d) Dehydration

| <b>PRACTICE:</b> Classify each one of the following reactions as phosphorylation (P), isomerization (I), or neither (N). |
|--|
| a) Conversion of glucose into glucose-6-phosphate.   |
| b) Conversion of glucose-6-phosphate into fructose-6-phosphate.  |
| c) Conversion of DHAP into G3P.  |
| d) Cleavage of fructose-1,6-bisphosphate into DHAP and G3P.  |
|  |
|  |
| <b>PRACTICE:</b> Which one of the following compound pairs is produced by cleavage of fructose-1,6-bisphosphate?         |
| a) Dihydroxyacetone phosphate and glyceraldehyde-3-phosphate   |
|  |

- b) glyceraldehyde-3-phosphate and glyceraldehyde-2-phosphate
- c) Glyceraldehyde-3-phosphate and CO<sub>2</sub>
- d) Dihydroxyacetone phosphate and 3-phosphoglycerate

PRACTICE: Identify which carbon atoms of fructose 1,6-bisphosphate become carbonyl groups in dihydroxyacetone phosphate and glyceraldehyde 3-phosphate when it undergoes a retro aldol reaction in step 4 of glycolysis.

- a) C3 and C4
- b) C2 and C5
- c) C3 and C5
- d) C2 and C4

**6** Oxidation × 2: G3P undergoes oxidation to produce 1,3-bisphosphoglycerate (1,3\_\_\_\_\_).

6a A hemithioacetal is formed first.

**6b** NAD+ \_\_\_\_\_ produces a thioester.

**6c** The thioester undergoes an \_\_\_\_\_ reaction with the phosphate ion.

7 Phosphate Transfer × 2: 1,3BPG produces 3-phosphoglycerate (3PG) by losing a \_\_\_\_\_ group.

□ Occurs via \_\_\_\_ mechanism.

- 8 Isomerization × 2: 3-phosphoglycerate (3PG) undergoes isomerization to yield 2-phosphoglycerate (2PG).
  - □ Occurs via \_\_ \_\_\_ reactions.

- 9 Dehydration × 2: 2-phosphoglycerate (2PG) undergoes dehydration to produce phosphoenolpyruvate (PEP).
  - (9a) Mg<sup>2+</sup> ions are required to reduce \_\_ charges.
- 9b Dehydration occurs via \_\_\_\_\_ mechanism.

- 10 Phosphate Transfer × 2: PEP yields pyruvate by losing its \_\_\_\_\_ group.
  - NAS reaction with ADP produces enolpyruvate.
- **10b** Tautomerization produces pyruvate.

| <b>EXAMPLE:</b> Which one of the fol | llowing alveolvsis | reactions will p | oroduce an ATP | molecule? |
|--------------------------------------|--------------------|------------------|----------------|-----------|
|--------------------------------------|--------------------|------------------|----------------|-----------|

- a) 3-Phosphoglycerate to 2-phosphoglycerate
- b) Glyceraldehyde-3-phosphate to 1,3-Bisphosphoglycerate
- c) Glucose to glucose-6-phosphate
- d) 1,3-Bisphosphoglycerate to 3-phosphoglycerate

**PRACTICE:** Identify the mechanism for each of the following reactions as NAS, E1cB, or S<sub>N</sub>2.

- a) \_\_\_\_\_ Dehydration of 2-phosphoglycerate to produce phosphoenolpyruvate.
- b) \_\_\_\_ Conversion of 1,3-bisphosphoglycerate into 3-phosphoglycerate.
- c) \_\_\_\_ Conversion of glucose into glucose-6-phosphate.
- d) \_\_\_\_ Conversion of 3-phosphoglycerate into 2-phosphoglycerate.

**PRACTICE:** If the P atom in 3-phosphoglycerate is labelled with phosphorous-32, where does the label end up after it is isomerized to 2-phosphoglycerate in step 8 of glycolysis?

- a) Ends up as the phosphate group in 2-phosphoglycerate.
- b) Ends up as free phosphate group because it is exchanged with a free phosphate from the solution outside the enzyme.
- c) On the histidine residue of the enzyme that catalyzes the reaction.
- d) Ends up as the y-phosphate group of ATP that's produced in the reaction.