

Endocrine Part 3

DIABETES MELLITUS

Pancreas.

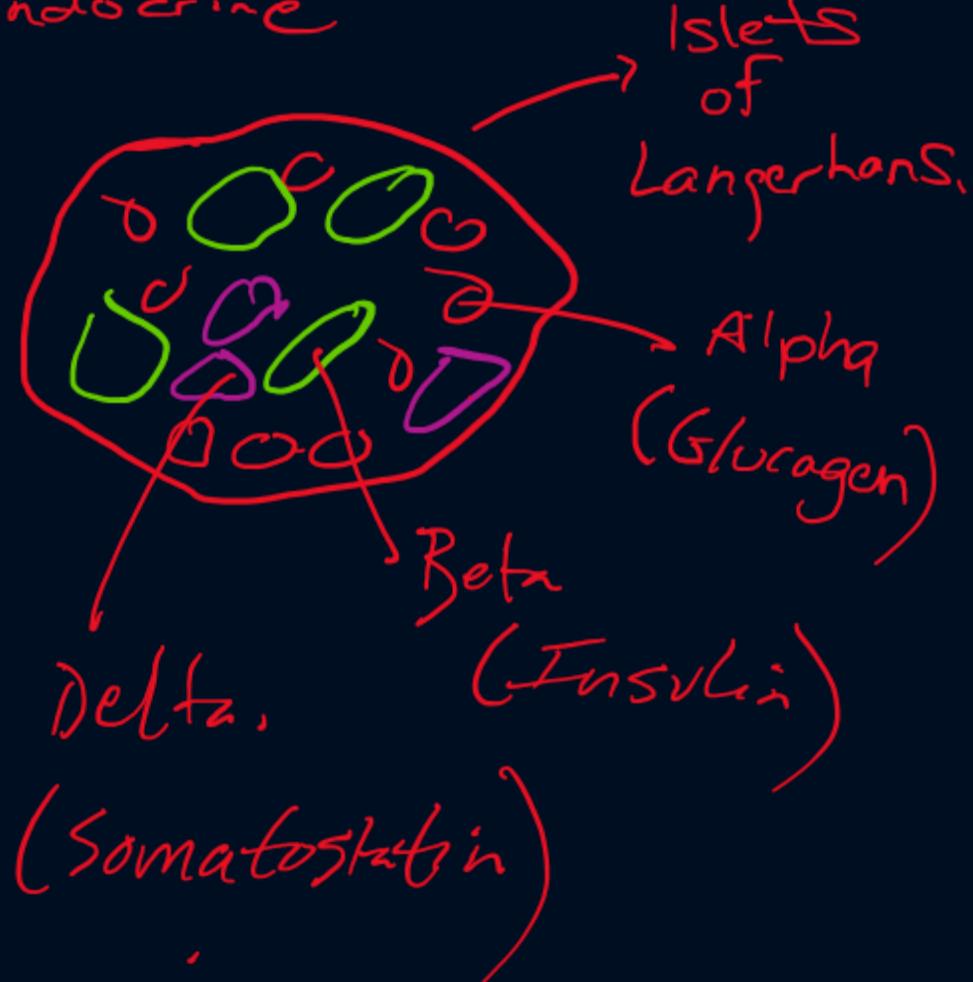


Homeostasis of
blood sugar.

\leftarrow $3.6 - 6.2 \text{ mmol/L}$
Hypoglycemia ($4 - 6$) \rightarrow

Hyperglycemia
(DKA; 10-15)

Endocrine



Glucose Metabolism.

Sandwich



Glucose.

} 2 ATP

(no O₂)

(O₂)

Aerobic
metabolism

Pyruvate

Anaerobic
metabolism

Lactic
Acid

Tetanol

Ketone
metabolism

↓

Acetyl CoA → fat
metabolism

↓

glycogen.

↓

Catab Cycle.

34 ATP

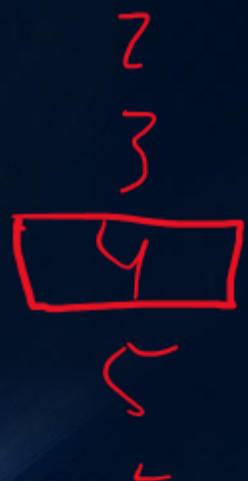
electron transport
chains

ATP (Walk-up stairs)

Glucose Metabolism and Regulation

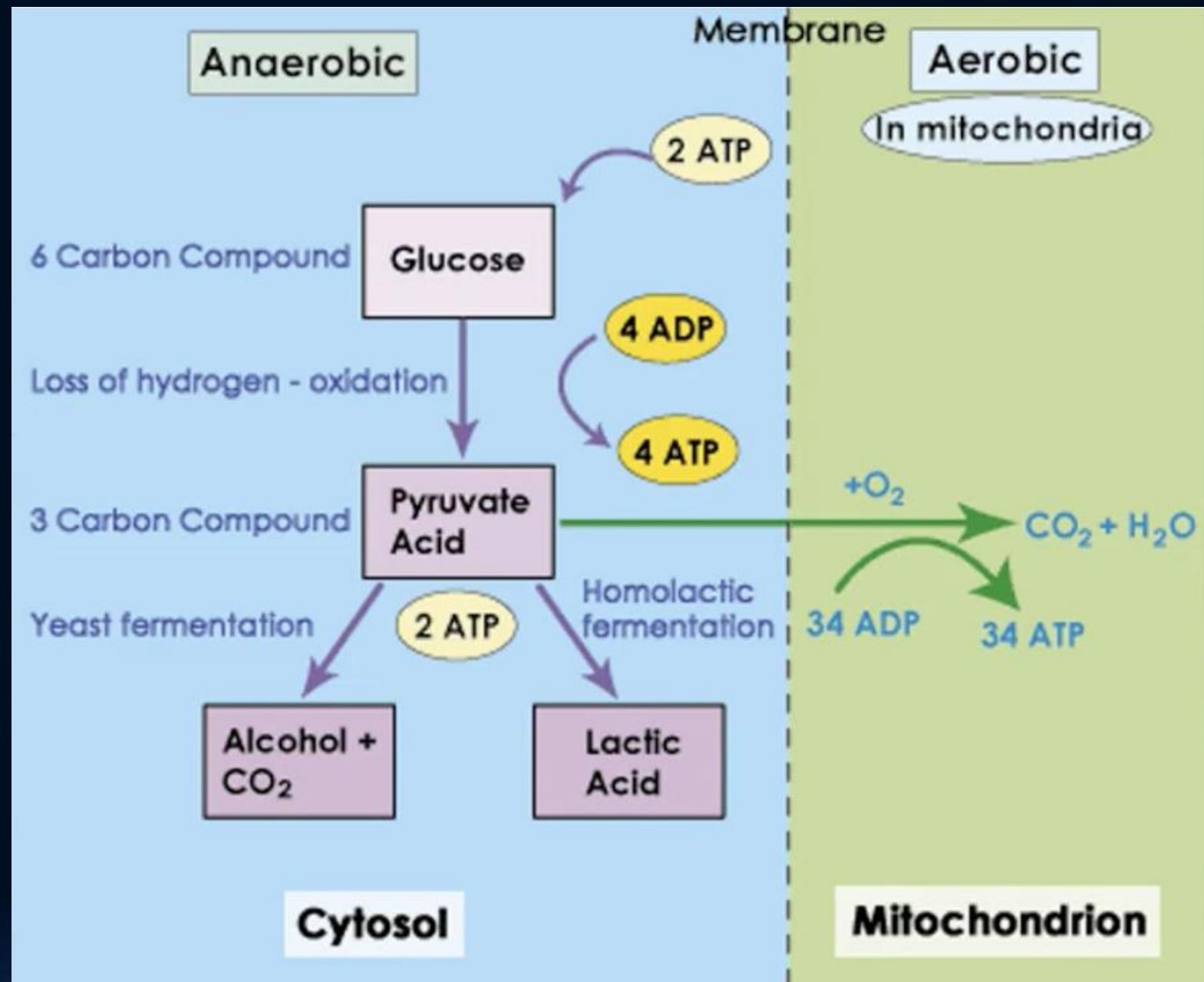
→ Glucose Transporters.

GLUT, 1 - 15



GLUT4 needs insulin

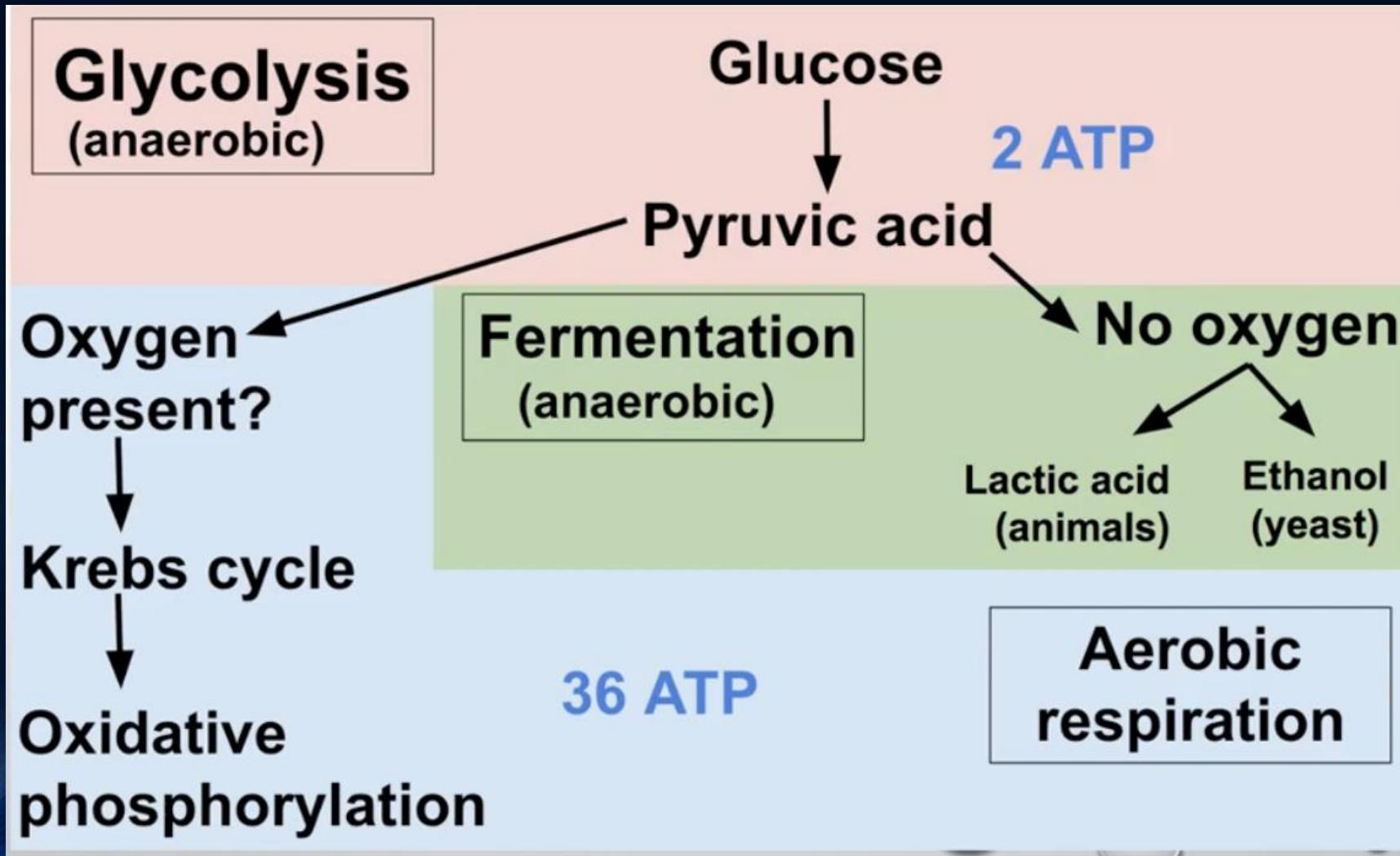
- Sk. muscle-
- Adipose tissue-



Glucose Metabolism

- In CHO metabolism, pyruvic acid (3C) is converted to Acetyl CoA (2C)
 - Acetyl CoA enters citric acid cycle in mitochondria to produce NADH and FADH₂
 - NADH and FADH₂ are used to make ATP during oxidative phosphorylation in the enzyme transport system
- In absence of glucose, cells will begin to metabolize lipids and then proteins
 - Lipids are chopped apart into 2C Acetyl CoA units which can enter into citric acid cycle; Acetyl CoA units can also combine to form acetoacetic acid – ketone bodies

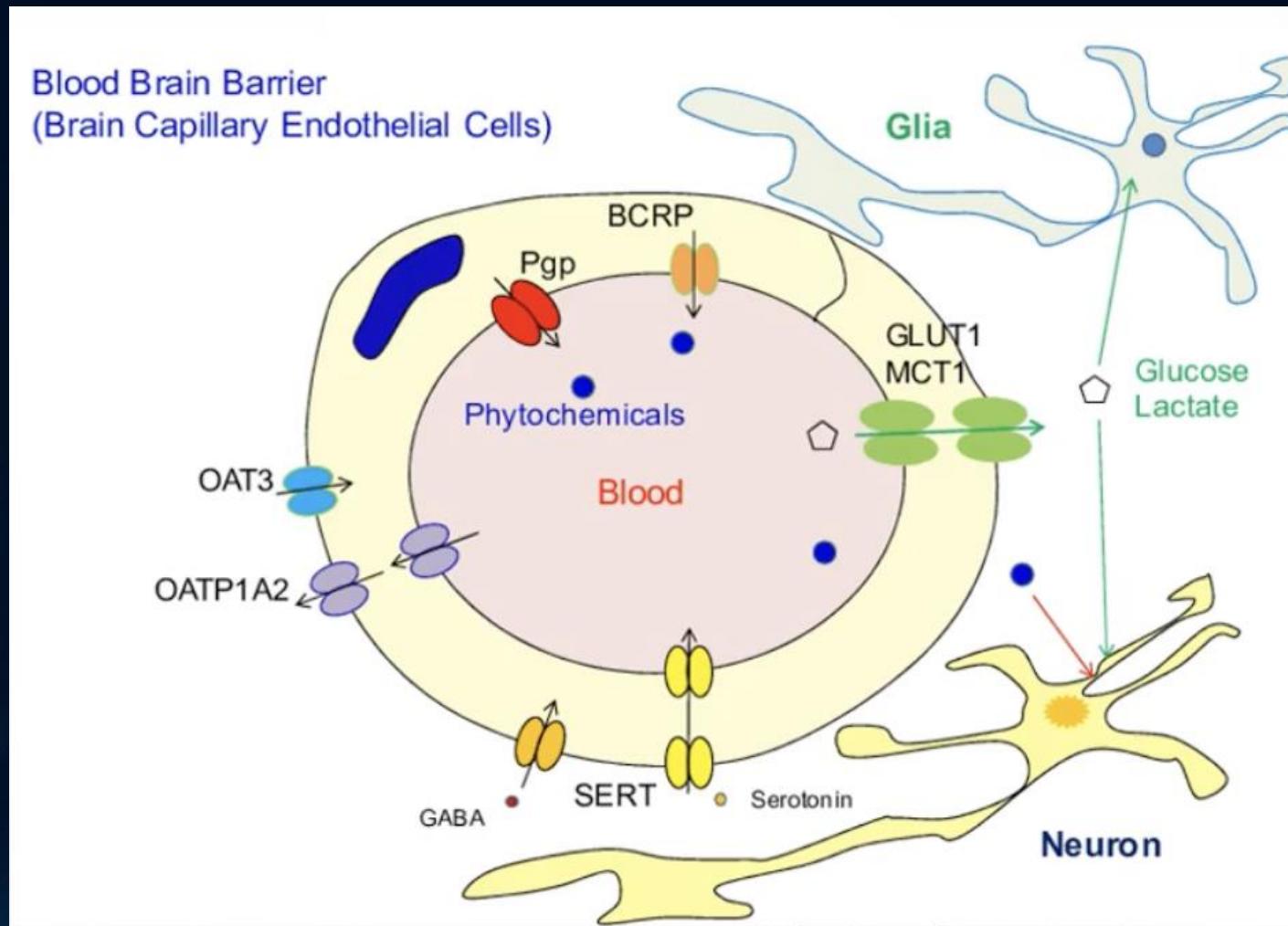
Glucose Metabolism



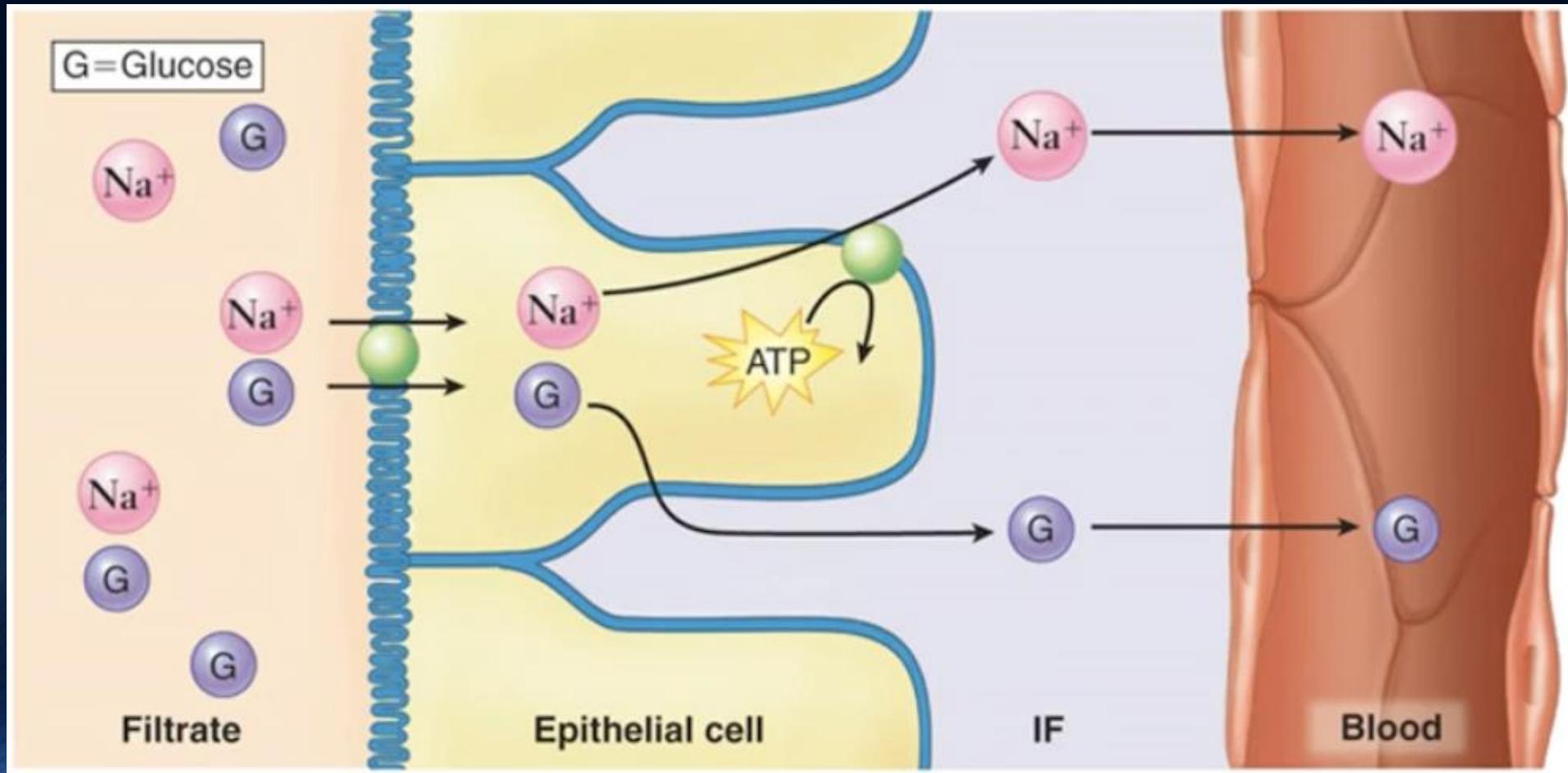
Glucose Transporters

- GLUT 1 – expression does not depend on insulin
 - Found in blood brain barrier – necessary for transporting glucose from the blood into the brain
 - Heart tissue
- GLUT 2 – expression does not depend on insulin
 - Found on liver cells
 - Pancreatic islet beta cells
 - Basolateral surface of tubule cells in kidney, brush border of GI
- GLUT 3 – expression does not depend on insulin
 - Nervous tissue
- GLUT 4 – expression depends on insulin
 - Skeletal muscle tissue
 - Adipose tissue

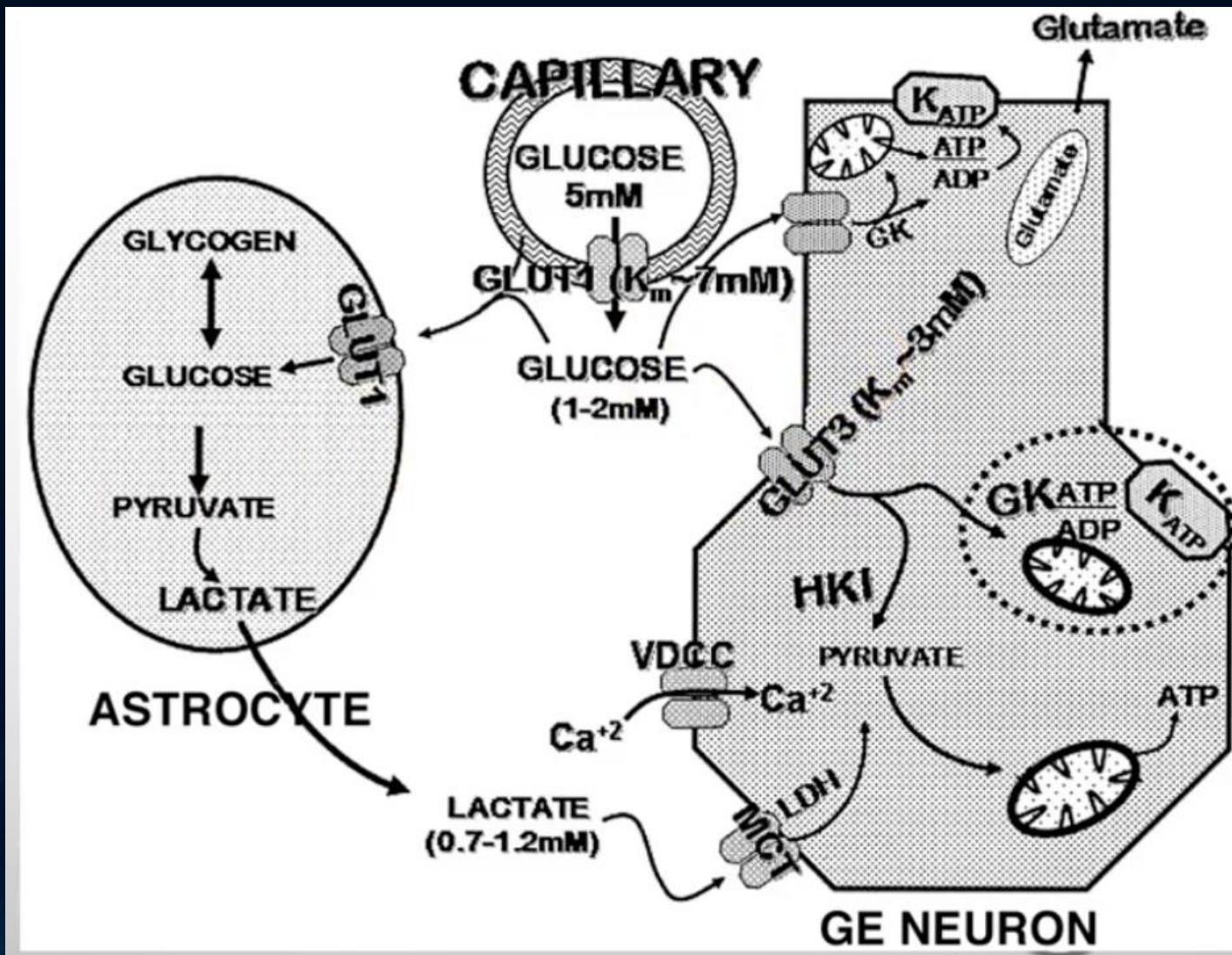
GLUT 1 Transporters



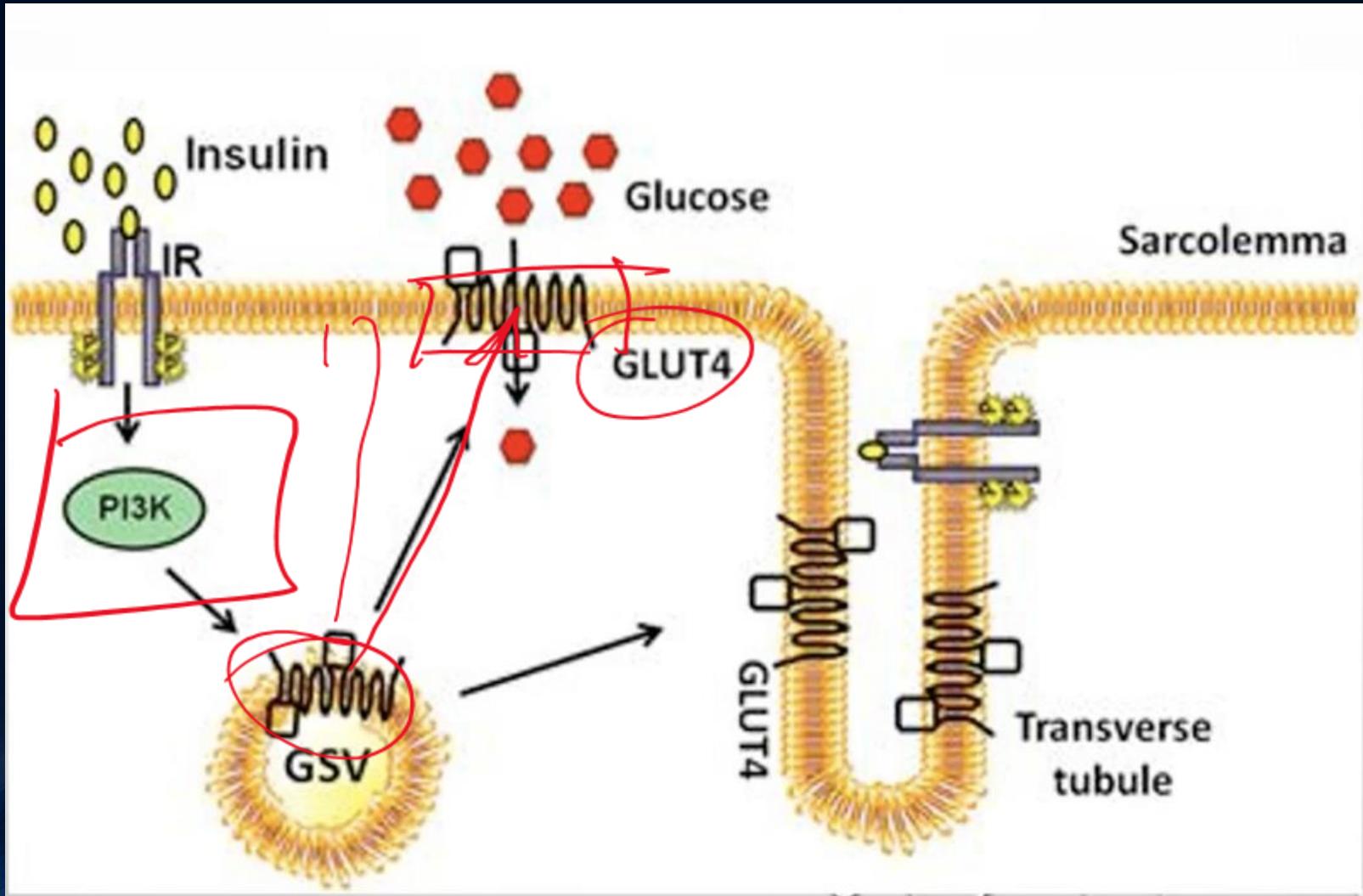
GLUT 2 Transporters



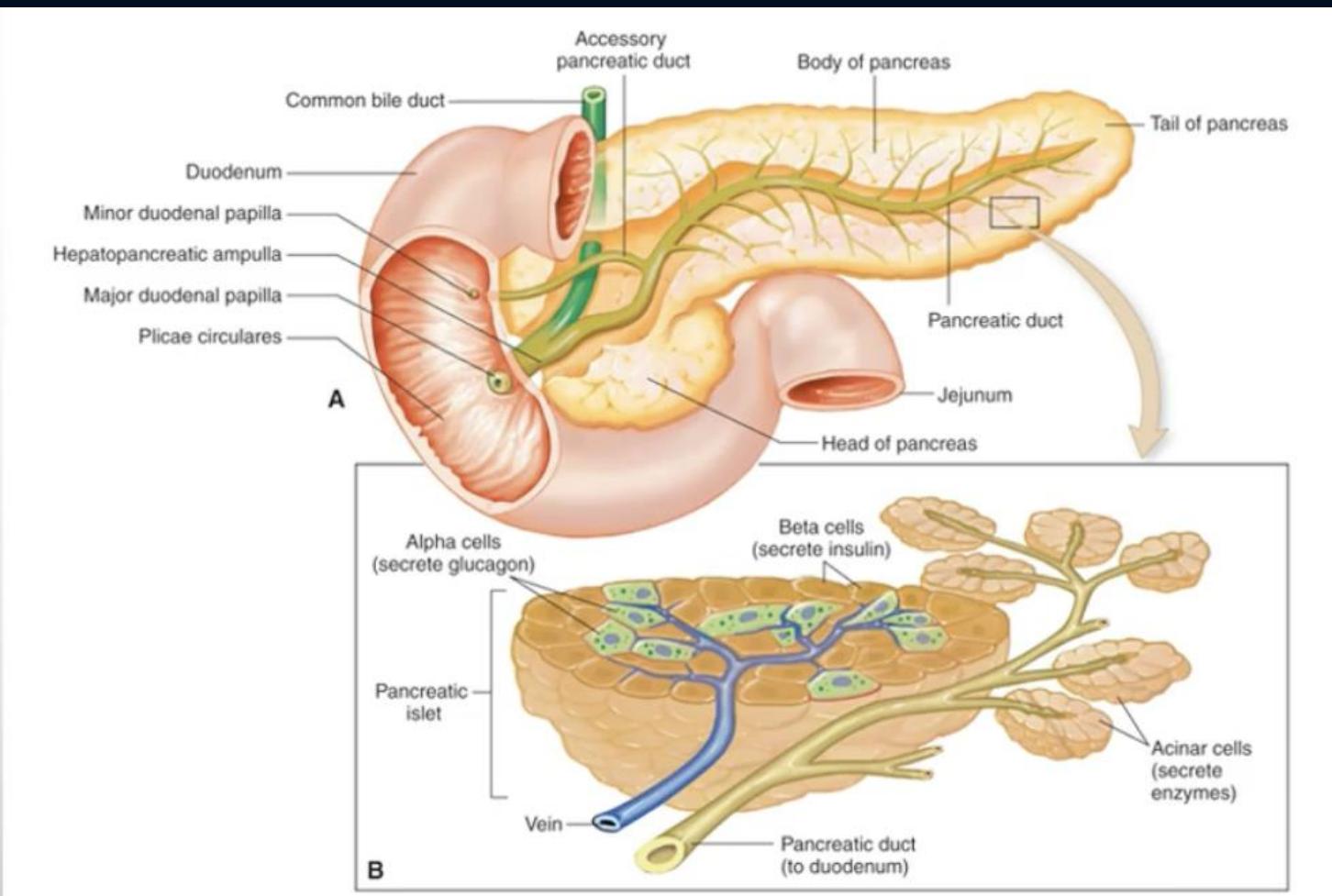
GLUT 3 Transporter

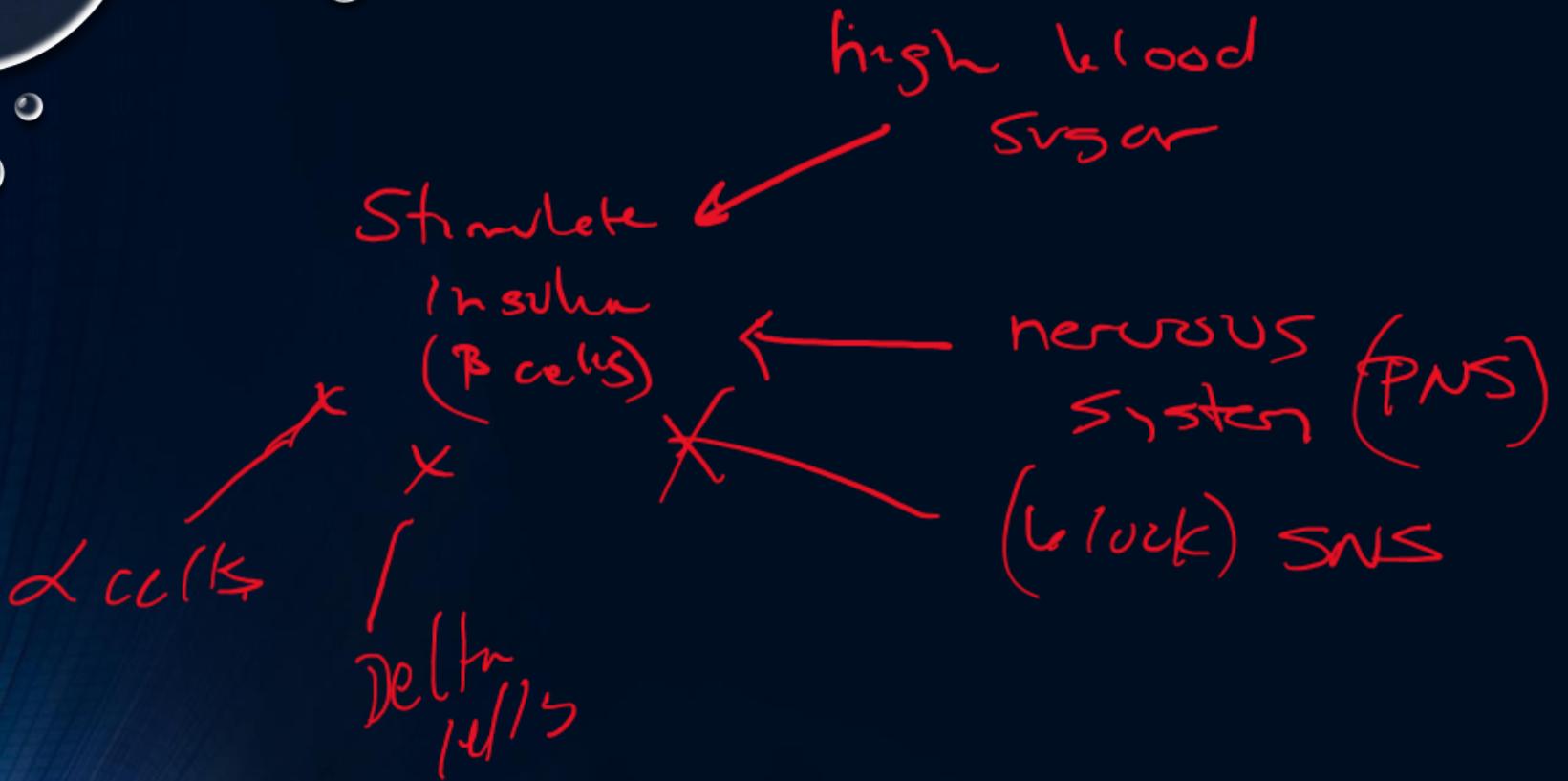


GLUT 4 Transporter



Role of Pancreas





Regulation of Blood Glucose - Insulin

- Stimulus for secretion by pancreatic beta islet cells ("fed state")
 - Increased blood glucose – sensed by glucokinase (enzyme that converts glucose to glucose-6-P)
 - Inactivation of PNS
- Secretion of incretins by small intestine
 - GIP, GLP-1
- Actions
 - Increases expression of GLUT4 transporters, especially on skeletal muscle cells, adipocytes
 - Promotes uptake of glucose into these cell types as well as liver cells
 - Promotes conversion of monomers – polymers (glycogen, TG, protein)
 - Net effect is to reduce blood glucose

Insulin (\downarrow blood Glucose)

→ bring glucose
into cells.

Fat production

Glycogenesis ↑

- liver
- Sk muscle.

Pr Fats

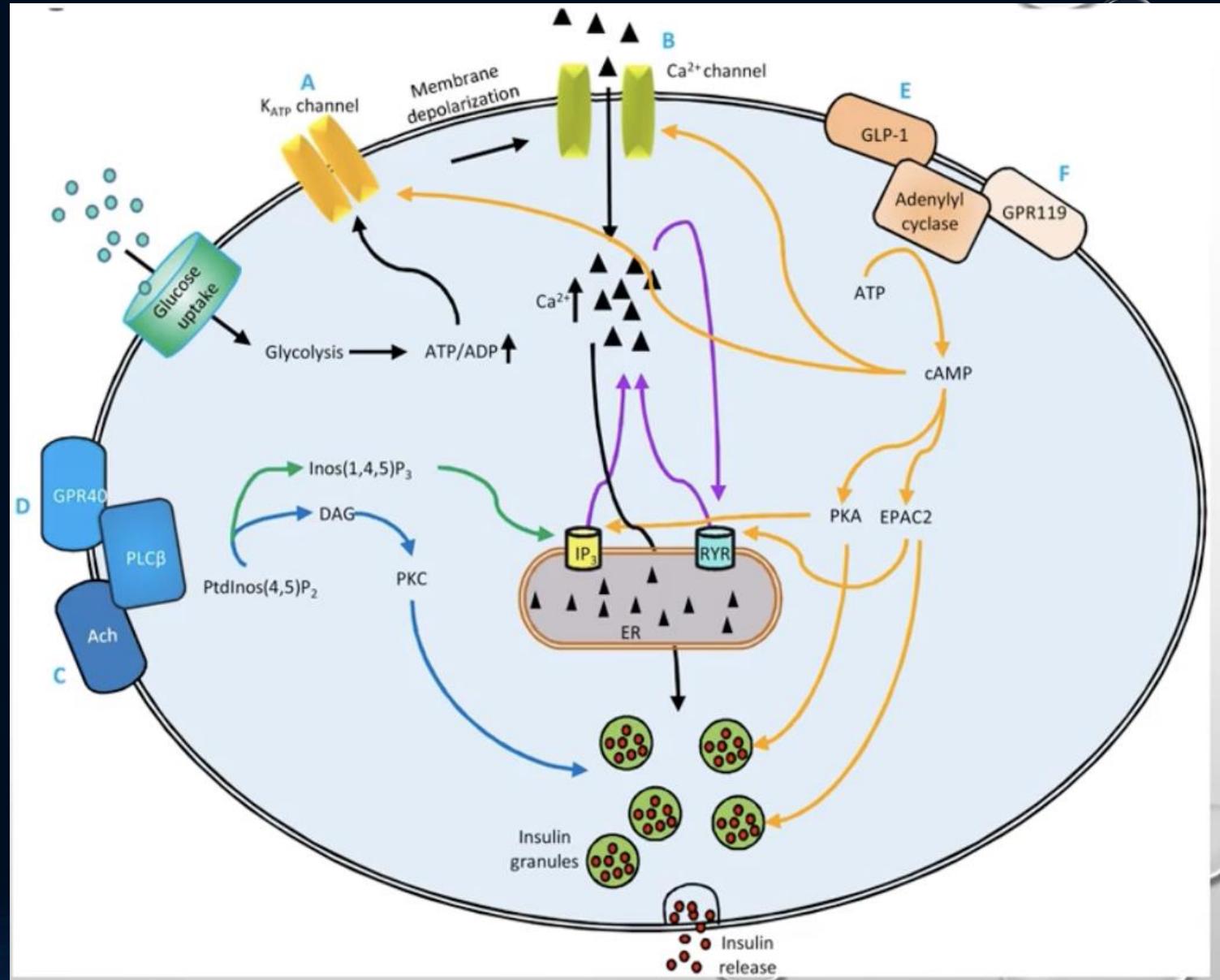
Gluconeogenesis

Glycogenolysis ↑

Glucagon:
 \uparrow blood sugar
(mobilizes stores)

Insulin Release

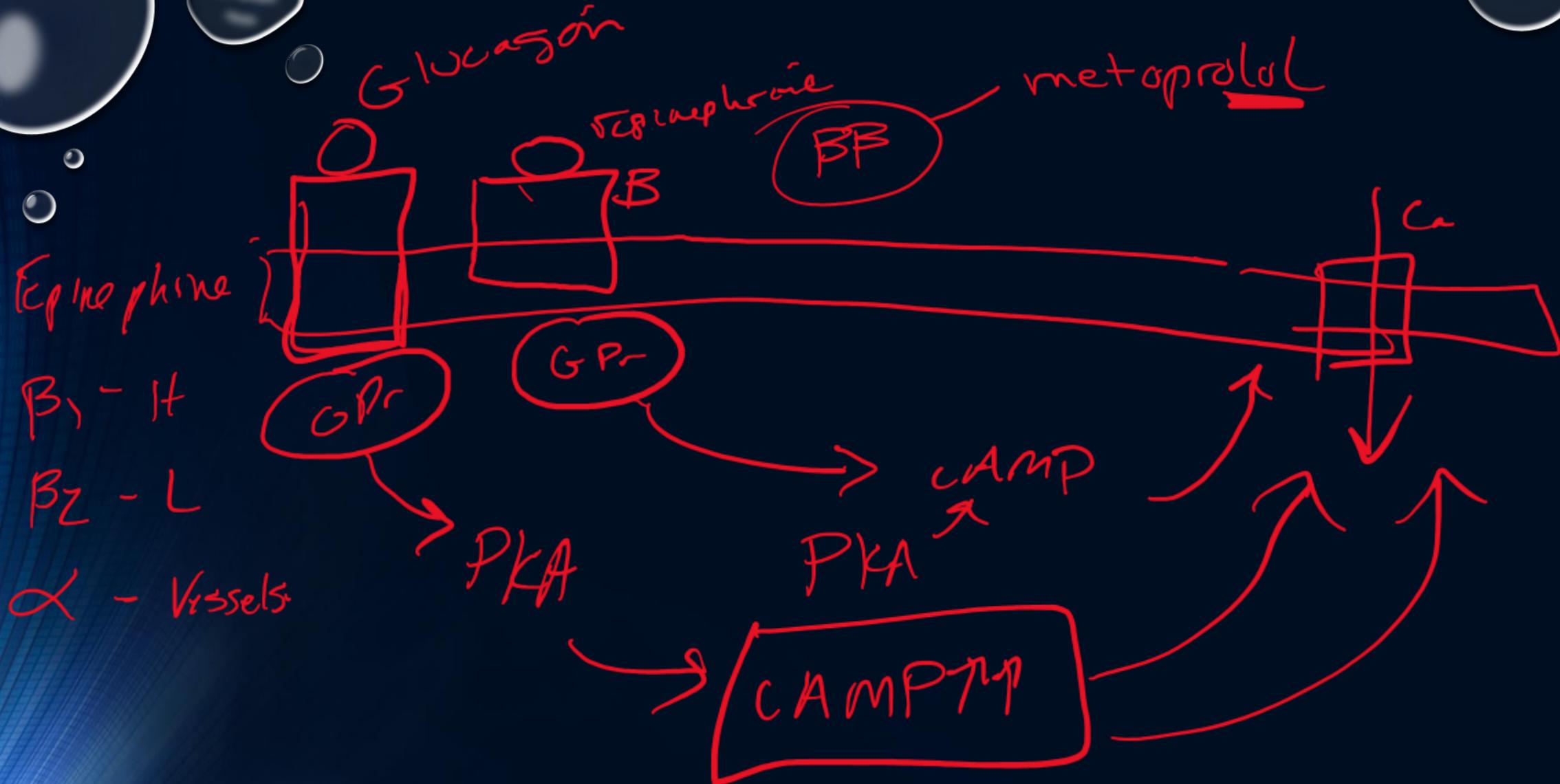
Structural classifications
of hormones



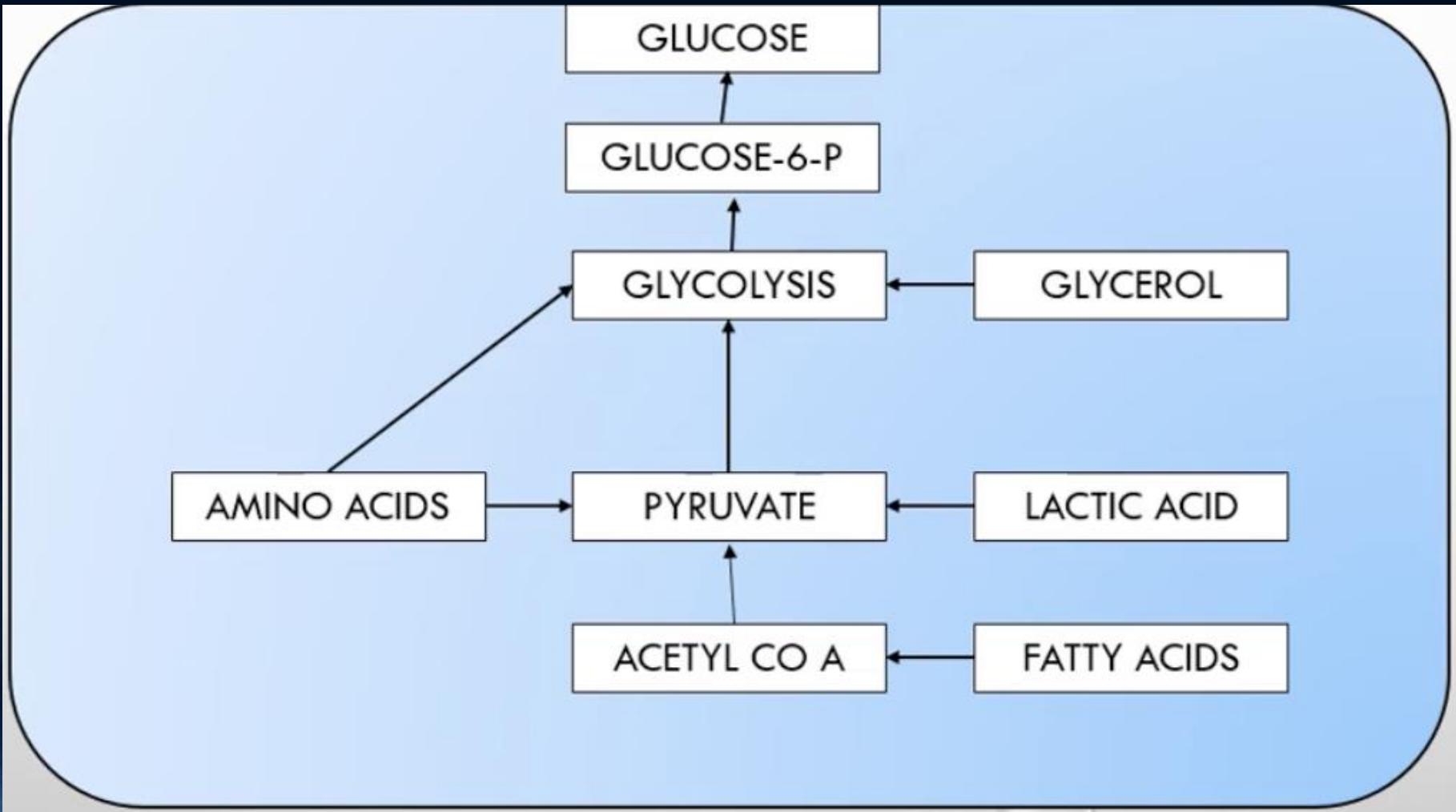
Regulation of Blood Glucose - Glucagon

- Stimulus for secretion by pancreatic alpha islet cells ("fasted state")
 - Decreased blood glucose
 - Increased activity of SNS via alpha-1 receptors
 - Glucagon secretion inhibited by incretins
- Actions – at liver cells, adipocytes and skeletal muscle tissue
 - Glycogenolysis: glycogen – glucose
 - Gluconeogenesis: AA/TG/Glycerol - glucose

→ $\beta\beta$) overdrive
→ Relaxation
Smooth muscle.
→ Anaphylaxis



Gluconeogenesis



Regulation of Blood Glucose - Others

- Epinephrine
 - Secreted by adrenal medulla in response to SNS activity
 - Effects
 - Directly stimulates release of glucagon
 - Stimulates skeletal muscle cells and liver cells – glycogenolysis / gluconeogenesis
- Glucocorticoids
 - Secreted by adrenal cortex following circadian rhythm, stress
 - Effects
 - Promotes gluconeogenesis by liver cells
- Growth Hormone
 - Secreted by anterior pituitary gland
 - Effects
 - Increased gluconeogenesis by adipocytes