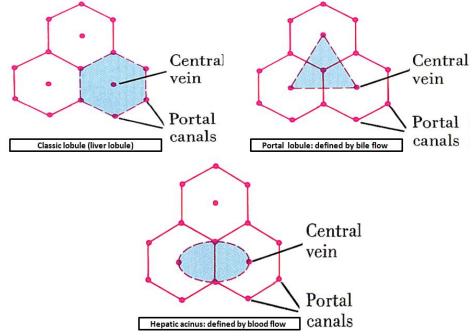
<u>Unit V – Problem 1 – Physiology of The Liver</u>

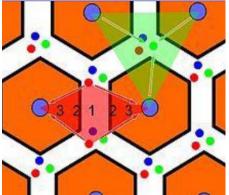


- What are some functions of the liver?

- Receiving blood from two sources (portal vein and hepatic artery) and serves as a blood reservoir (450 ml to 1 L).
- Cleaning blood via liver macrophages (Kupffer cells).
- Production of 50% of lymph.
- The liver regenerates at a high rate.
- Morphofunctional element:



- Liver structure (structure of the lobule): there are three zones
 - **Periportal zone** (1): oxidative liver functions
 - ✓ Gluconeogenesis.
 - ✓ Ureagenesis.
 - ✓ β-oxidation of fatty acids.
 - \checkmark Cholesterol synthesis.
 - ✓ Bile formation.
 - Transition zone (2).
 - Centrilobular zone (3):
 - ✓ Glycolysis.
 - ✓ Glycogenesis.
 - ✓ Lipogenesis.
 - ✓ Ketogenesis.
 - ✓ Glutamine formation.
 - ✓ CYP-450 based drug detoxification.

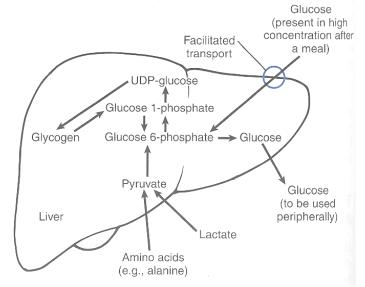


- Blood flow in liver lobule:

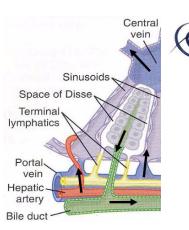
- The liver gets 65% of total oxygen supply from hepatic artery.
- Blood is flowing from portal vein and hepatic artery to sinusoids and then to the central vein which drains into the inferior vena cava.
- Pressure gradients (as blood is flowing toward the central vein):
 - ✓ 80-120 mmHg in hepatic artery.
 - ✓ 3-5 mmHg in sinusoids.
 - ✓ 1-3 mmHg in central vein.

Liver functions:

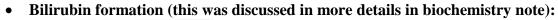
- **Carbohydrate metabolism**: glucose buffer, glycogenesis, glycogenolysis and gluconeogenesis (from lactate, glycerol and amino acids).
 - ✓ After having a meal, glucose will be present in high level in blood → transported to the liver via active Na⁺-glucose co-transport → glucokinase will add a phosphate group to the 6th carbon position in glucose molecule to generate glucose-6-phosphate which can be converted to UDP-glucose and then used to synthesize glycogen.
 - ✓ Lactate is generated from glycogen breakdown in muscles and then transported to liver to be converted into pyruvate and then used in gluconeogenesis (which is stimulated by glucagon & NE and inhibited by insulin).

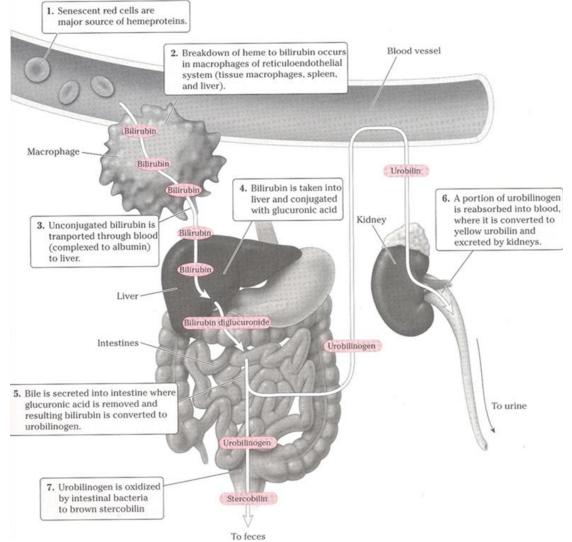


- **Fat metabolism**: β-oxidation of fatty acids; cholesterol, phospholipid and lipoprotein synthesis; synthesis of triglycerides
 - ✓ β-oxidation generates acetyl-CoA → if present in excess → acetyl CoA is used to generate ketone bodies:
 - ✤ Acetoacetic acid.
 - * β-hydroxybutyric acid.
 - ✤ Acetone.
 - ✓ In the liver, fatty acids are used to synthesize triglycerides which will be packed into VLDL and then transported to adipose tissue to be stored there.
- In the liver, amino acids will enter the urea cycle and will be used in the synthesis of plasma proteins.
- Bile secretion and storage:
 - ✓ 600-1000 ml is produced per day.
 - \checkmark Bile acids serve as emulsifiers and form micelles.
 - \checkmark Bile is used for excess cholesterol excretion.



- ✓ <u>Bile is composed of:</u>
 - ↔ Water.
 - ✤ Bile salts.
 - ✤ Bilirubin.
 - ✤ Cholesterol.
 - ✤ Fatty acids.
 - ✤ Lecithin.





- Coagulation proteins:
 - Fibrinogen; prothrombin; factors V, VII, IX, X, XI; proteins C and S; antithrombin.
- Detoxification and degradation of:
 - ✓ Drugs (CYP-450 family).
 - ✓ Hormones: thyroxine, steroids (estrogen, aldosterone and cortisol).
 - ✓ Ca^{2+} excretion.
 - ✓ Activation of vitamin D (with kidneys).
 - ✓ Synthesis of somatomedins.

