

## **Biochemical Aspects of Bile Acids and Salts**



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## **Objectives**

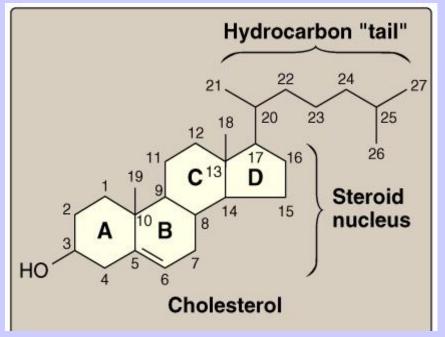
- > Structure of primary bile acids and salts
- Structure of secondary bile acids and salts
- **Functions of bile salts**
- Enterohepatic circulation
- > Malabsorption syndrome
- Cholelithiasis

## **Cholesterol**

**Cholesterol** (27 C) is the:

Parent steroid compound

**Precursor of bile acids and salts** 



## **Primary Bile Acids**

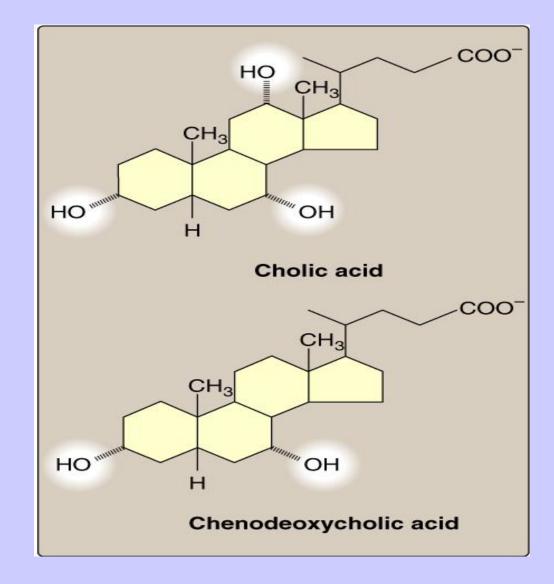
Primary bile acids (24 C):

Amphipathic

-COOH at side chain

Cholic acid: 3 OH

**Chenodeoxycholic: 2 OH** 



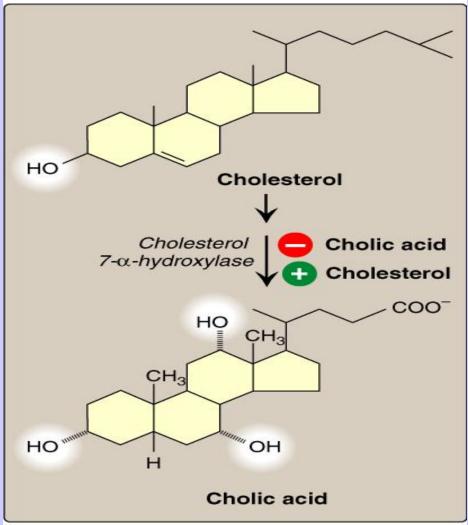
## **Hepatic Synthesis of Bile Acids**

The rate-limiting step is catalyzed by:

**Cholesterol 7-α-hydroxylase** 

**Regulation: Down-regulated by end products** (bile acids) "Enzyme repression"

**Up-regulated by cholesterol "Enzyme induction"** 



## **Primary Bile Acids and Salts**

# Cholic acidBILE ACIDSChenodeoxycholic acidGlycocholicBILE SALTSGlycochenodeoxycholicTaurocholicBILE SALTSTaurochenodeoxycholic

Bile salts (Conjugated bile acids): amide-linked with glycine or taurine The ratio of glycine to taurine forms in the bile is 3:1

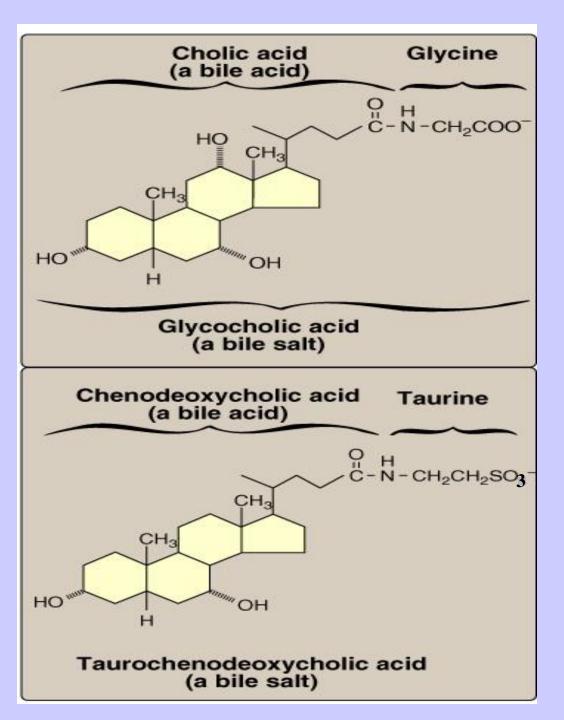
## **Bile Salts**

- Addition of glycine or taurine results in the presence of fully ionized groups at pH 7.0:
  - -COO<sup>-</sup> of glycine &
  - -SO<sub>3</sub><sup>-</sup> of taurine
  - (hence, its name as bile salts e.g.,
  - Sodium or potassium glycocholate)
- > More effective detergent than bile acids
- > Only bile salts, but not acids, found in bile

## **Bile Salts**

Na or K Glycocholate

Na or K Taurochenodeoxycholate



# Hormonal Control of Bile Secretion

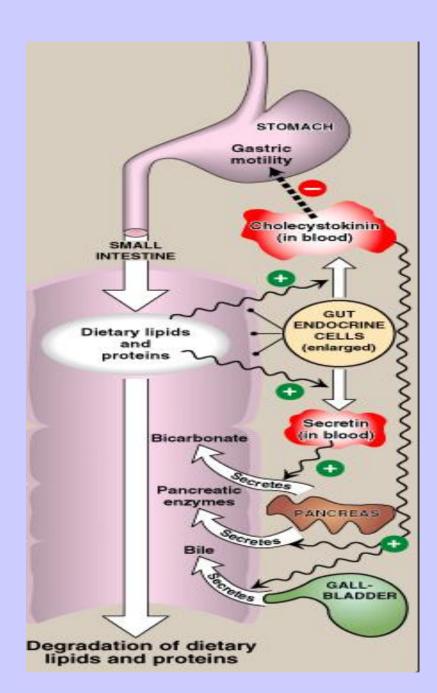
#### **Stimulus:**

Undigested lipids and partially digested proteins in duodenum

#### Hormone from gut cells: Cholecystokinin (CCK)

#### **Responses:**

- 1. Secretion of pancreatic enzymes
- 2. Bile secretion
- 3. Slow release of gastric contents



## **Functions of Bile Salts**

- Important for cholesterol excretion:
  - **1. As metabolic products of cholesterol**
  - 2. Solubilizer of cholesterol in bile
- Emulsifying factors for dietary lipids, a prerequisite step for efficient lipid digestion
- Cofactor for pancreatic lipase and PLA2
- Facilitate intestinal lipid absorption by formation of mixed micelle

## **Emulsification of Dietary Lipids in Duodenum: Role of Bile Salts**

- Emulsification increases the surface area of lipid droplets, therefore the digestive enzymes can effectively act.
- Mechanisms:
  - **1. Mechanical mixing by peristalsis**
  - 2. Detergent effect of bile salts:

Bile salts interact with lipid particles and aqueous duodenal contents, stabilizing the particles as they become smaller, and preventing them from coalescing. Absorption of Lipids by Intestinal Nucosal Cells: Role of Bile salts Mixed micelles: Disc-shaped clusters of amphipathic lipids. Arranged with their hydrophobic groups on the

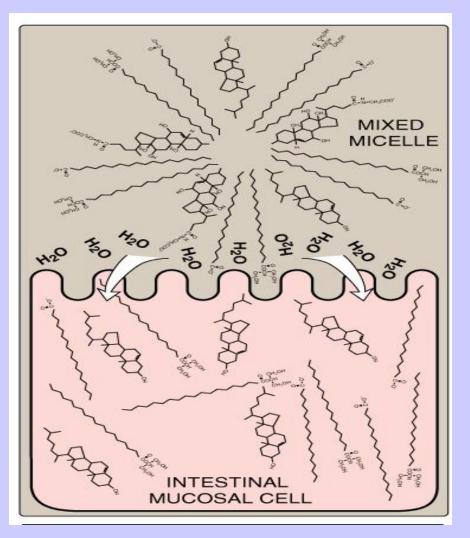
inside and their hydrophilic groups on the outside.

Micelle includes end products of lipid digestion, bile salts and fat-soluble vitamins

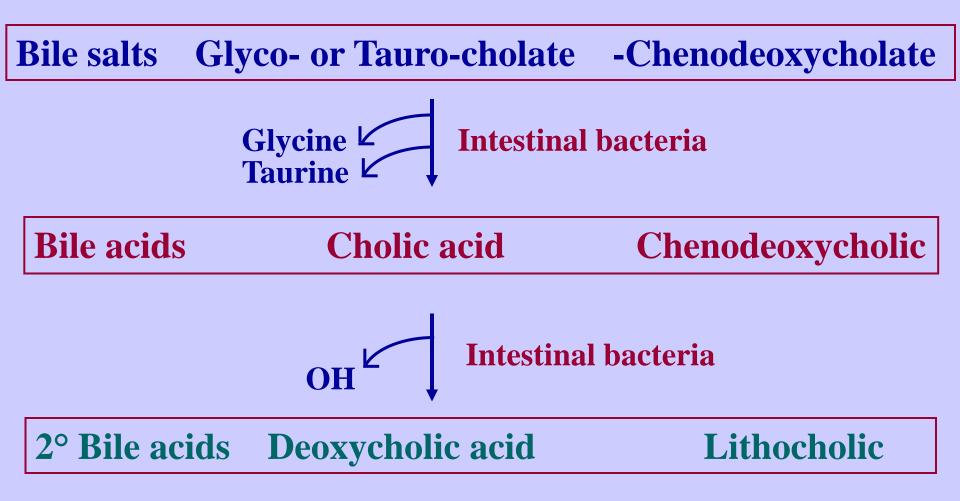
**Note:** Short- and medium-chain fatty acids do not require mixed micelle for absorption by intestinal cells

## **The Role of Bile Salts in Absorption of Lipids by Intestinal Cells**

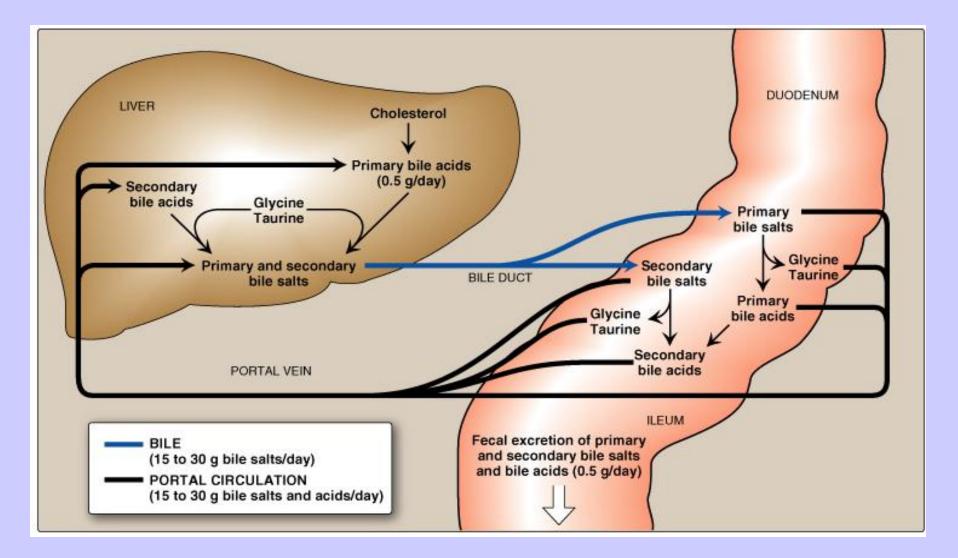
Mixed Micelle Formation: Bile salts End products of lipid digestion Fat-soluble vitamins



## **Secondary Bile Acids**



## **Enterohepatic Circulation**

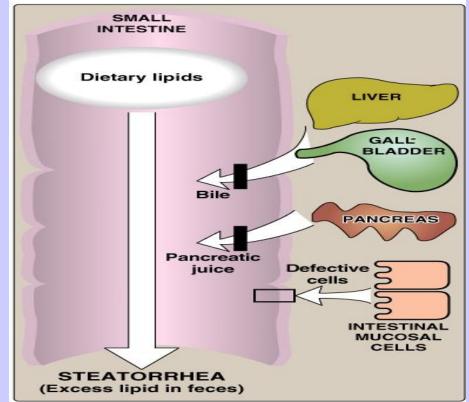


## Maldigestion/Malabsorption of Lipids

#### **Decreased bile secretion by:**

Liver diseases: e.g., Hepatitis or cirrhosis

Gall bladder diseases: e.g., Gall stones



## **Cholelithiasis**

### **Causes:**

- ↓ Bile salts in bile: Biliary tract obstruction (intereferes with enterohepatic circulation) Hepatic dysfunction (↓ synthesis)
- **†** Biliary cholesterol excretion

## **Treatment:**

Bile acid replacement therapy
Surgical

## **THANK YOU ③**