# Lipid Metabolism

Course Code: ZOOL 4008 (Biochemistry and Metabolism) M.Sc. (Zoology), Semester –II



## lipid bilayer

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### Dr. Shyam Babu Prasad

Assistant Professor Department of Zoology Mahatma Gandhi Central University (MGCU), Motihari-845401 (Bihar) Email: shyambabuprasad@mgcub.ac.in

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phospholipid molecule

# Lipids

•A lipids are non-polar organic compounds that is insoluble in water and soluble in alcohol, ether, and chloroform.

•Lipids are an important constituent of the structure of living cells. Together with carbohydrates and proteins, lipids are the main constituents of plant and animal cells.

•Lipids are easily stored in the body and serve as a source of Energy/Fuel.

•Lipids includes fatty acids, neutral fats, waxes and steroids (like cortisone) etc. Almost all fat in your diet comes in the form of triglycerides

•Fatty acids rarely occur as free molecules in nature. it is found in complex molecules such as fats (energy-storage compounds) and phospholipids (the primary lipid components of cellular membranes).

# Lipids Classification

#### Simple Lipids:

Esters of fatty acids with various alcohols. They are Saturated and Unsaturated (Ex. Fats, oil and Waxes)

#### **Complex Lipids:**

Esters of fatty acids with various alcohols + Other Groups like phosphoric acids, sugar residue etc. for ex.

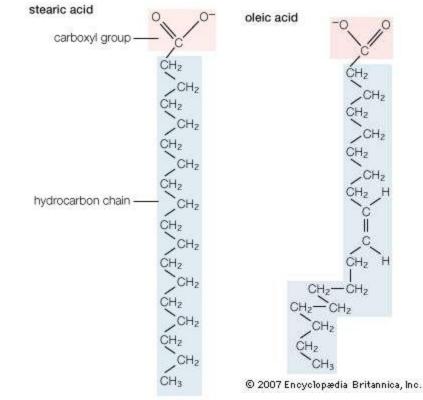
- A. Phospholipids (glycerophospholipid s and sphingophospholipid s )
- B. Glycolipids (glycosphingolipids) : Carbohydrates containing lipids.
- C. Other complex lipids: Sulfolipids, aminolipids, and Lipoproteins.

# Precursor and derived Lipids:

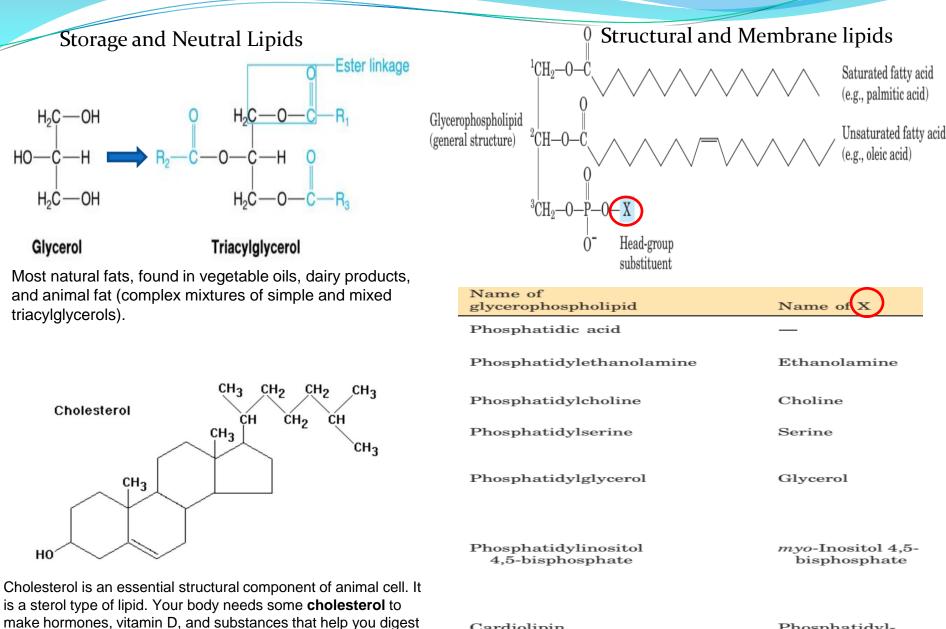
Steroids, other alcohols, fatty aldehydes, and ketone bodies, lipid-soluble vitamins, and hormones.

## **Structure of Lipids**

- Fatty acids are composed of a <u>hydrocarbon</u> chain with one terminal carboxyl group (COOH). At physiological condition hydrogen ion (H<sup>+</sup>) lost to form a negatively charged carboxylate group (COO<sup>-</sup>).
- Most biological fatty acids contain an even number of <u>carbon</u> atoms, because the biosynthetic pathway is common to all organisms.
- Fatty acids are amphipathic in nature.
- They are found in two form saturated (only single bond), Ex. myristic acid (C14), palmitic acid (C16), Stearic acid (C18), etc. and Unsaturated (double bond), Ex. palmitoleic acid (C16), Oleic acids (C18) etc.



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foods

Cardiolipin

Phosphatidylglycerol

## Lipid Metabolism

•Lipid metabolism is referred to the synthesis and degradation of lipids within the cells, either break down or storage of fats for energy.

•These fats are obtained from consuming food and absorbing them or they are synthesized by an animal's liver.

•Minimal amount of fat is essential in our food :
✓ For essential fatty acids synthesis .
✓ Help Fat-soluble vitamins absorption.
✓ As a Source of energy: 1 gm supplies 9.1 calories

• Just like glucose Metabolism, the end-products of fatty acid metabolism are carbon dioxide, water and ATP.

•However, complete combustion of fatty acids requires glucose to convert it in to carbon dioxide, water and ATP, otherwise ketones are produced.

Digestion and Absorption of the Lipids

There are following steps that start from mouth:

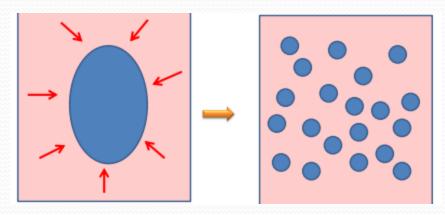
#### Table 1. Steps of lipid digestion and absorption

Step	Location	Enzymes
1. Minor digestion (TAGs $\rightarrow$ DAGs + FFA)	mouth and stomach	lingual/gastric lipase
2. Major digestion (all) TAG $\rightarrow$ MAG + 2FFA (PL) CE $\rightarrow$ chol. + ester (CE) PL $\rightarrow$ FA + lysoPL (PLA)	lumen of the small intestines	pancreatic lipase (+colipase) cholesterol esterase phospholipase A <sub>2</sub>
3. Formation of mixed micelles (uses bile salts as biological detergent)	lumen of the small intestines	N/A
4. Passive absorption of lipolytic products	into intestinal epithelial cell	N/A
5. Assembly and export of chylomicrons	from intestinal cells to the lymphatics	N/A

## 1. Digestion of Triglycerides

#### It is a two steps process:

**A. Emulsification**: Breakdown of large fat globule smaller ones

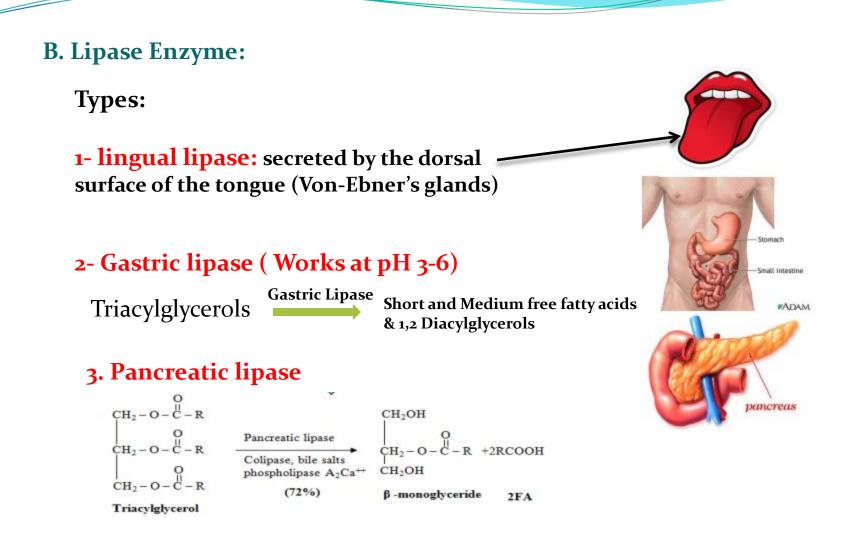


#### It Occurs in:

✓ Mouth by chewing.

Stomach by peristaltic contractions.

✓ Intestine by peristaltic movement, bile salts, lysophospholipids



**4. Intestinal Lipase:** Act within intestinal mucosal cells, it hydrolyse the absorbed primary ( $\alpha$ ) monoglycerides forming glycerol and FFA,

Note# The most active is pancreatic lipase

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2. Digestion of Phospholipids:

Phospholipase Phospholipase Iysophospholipids

•Intestinal phospholipase may complete the hydrolysis of lysophospholipids.

•Phospholipase A2 activated by trypsin & requires bile salts for activity

3. Digestion of Cholesterol esters:

Cholesterol esters FA & free cholesterol

**Absorption of Lipids** 

The end products of lipid digestion are: monoglycerides, FA, glycerol, cholesterol & lysophospholipids.

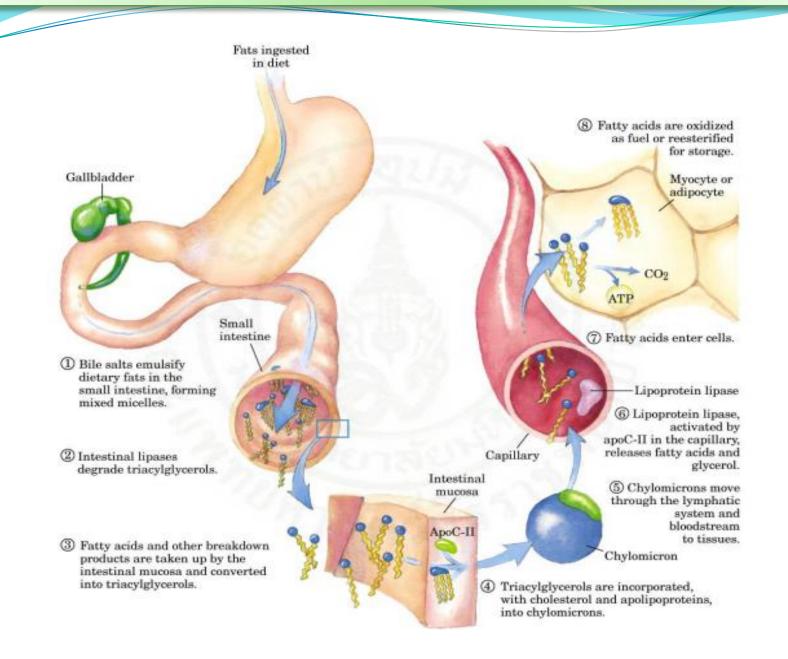
**1- Glycerol and short chain FA :** 

Water soluble *matheful* carried through portal circulation

2- Long chain FA, monoglycerides, cholesterol & lysophospholipids: Need bile salts to be absorbed

Note # Immediately after absorption of lipids there is turbidity in plasma observed due to circulating chylomicrons (appear in plasma up to 2 hours after meals). •Chylomirons are ultra low-density lipoproteins (ULDL), that consist of triglycerides (85–92%), phospholipids (6–12%), cholesterol (1–3%), and proteins (1–2%). • This turbidity is soon cleared by lipoprotein lipase enzyme (clearing factor). Lipoprotein lipase breaks Chilomicrons into Glycerol and FA.

## **Mechanism of Lipids transport and Absorptions**



# **Fatty Acid Synthesis**

• Fatty acids Synthesis start in cytoplasm with help of acetyl-CoA and NADPH produced from mitochondria and Pentose phosphate Pathway (PPP) respectively using enzyme fatty acid synthases.

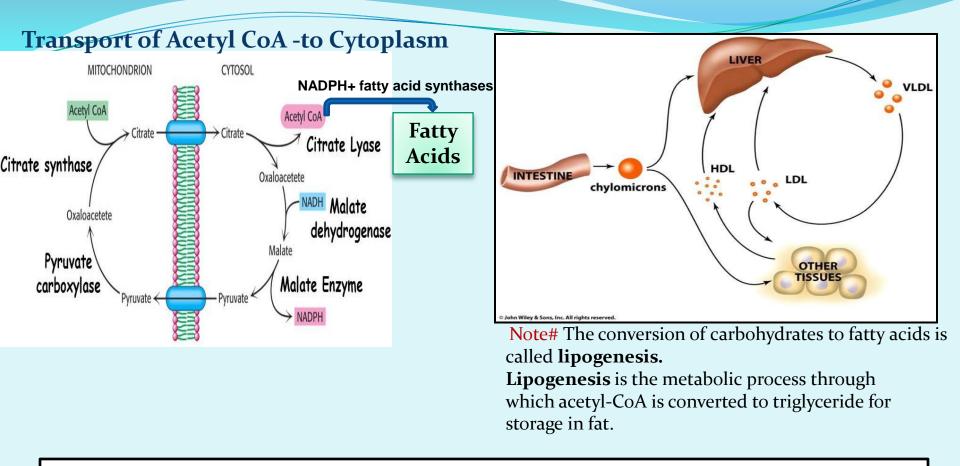
• First acetyl-CoA transported from mitochondria **through citrate—malatepyruvate –Shuttle** in form of citrate, which further breaks into Acetyl-CoA and Oxaloacetate.

• Only small chain fatty acids get synthesized in to cytoplasm, so the synthesis of long chain fatty acids like Triacylglycerol required specialized organ like Liver.

•The liver is the major site for converting excess carbohydrates and proteins into fatty acids and triglyceride. The liver synthesizes large quantities of **cholesterol and phospholipids**.

• After synthesis, VLDL lipoproteins are then exported through blood circulation and stored in adipose tissue.

• A small fraction is also converted to small ketone molecules that are exported via the circulation to peripheral tissues, where they are metabolized to yield energy .



VLDL (very low-density lipoprotein): Bad" cholesterol because it contributes to the build up of plaque in arteries). It acts as transport vehicles for triglycerides. LDL (low density lipoproteins): It is also called bad" cholesterol. It carry about 75% total blood cholesterol.

HDL (High-density lipoprotein): It is called good cholesterol because remove excess cholesterol, prevent plaque formation in arteries and protect from heart Diseases.

## Important Functions of Lipids

- ✓ As an energy source, lipids provide 9.1 Kcal of energy per gram.
- ✓ Triglycerides provide energy storage in adipocytes.
- ✓ Phosphoglycerides, sphingolipids and steroids are structural component of cell membrane .
- ✓ Steroids hormones are critical intracellular messengers.
- ✓ Lipid –soluble vitamins (A,E,D,K).
- ✓ As protective form of the cells of many bacteria, leaves of higher
- plants, exoskeleton of insects and the skin of vertebrates.
- ✓ Act as regulatory substances.
- ✓ As transport form of some neurotransmitters.
- ✓ As receptors in nerve ending membranes.
- ✓ As determinants of immunological specificity.
- ✓ Enzyme co-factors

References:

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• Lehninger principles of biochemistry (4th ed.): Nelson, D., and Cox, M, W.H. Freeman and Company, New York, 2005. Thank You