



Biological Chemistry Department

Biological Chemistry

CONJUGATED PROTEINS

Speciality: Pharmacy for foreign students (Language of instructions - English)

Lecturer: ass. prof. Kravchenko G.B.



Lecture Plan


1. Conjugated proteins classification.
2. Chromoproteins.
 - 2.1. Hemoproteins, structure and functions.
3. Glycoproteins and proteoglycans, structure and functions.
4. Lipoproteins, structure and functions.
5. Metaloproteins, structure and functions.
6. Phosphoproteins, structure and functions.

Individual work

1. Glycoproteins as pharmaceutical preparations.

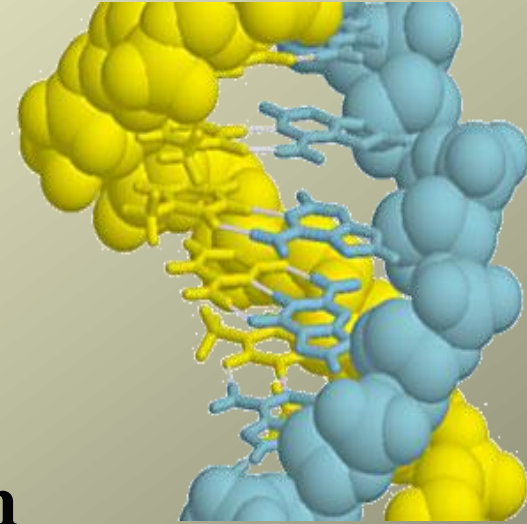


Information Resources

1. *Biological Chemistry: Textbook* / A.L. Zagayko, L.M. Voronina, G.B. Kravchenko, K.V. Strel`chenko. - Kharkiv: NUPh; Original, 2011. - 33-47 p.
 2. *Training Journal for Licensed Exam "KROK-1": Study Material in Biological Chemistry.* - Kharkiv: NUPh, 2017. - 18-26 p.
 3. *Laboratory Manual on Biochemistry.* Kharkiv: NUPh, 2017. - 29-31 p.
 4. *Hemoglobin and Myoglobin: The Medical Biochemistry Page.* Available on:
<https://themedicalbiochemistrypage.org/hemoglobin-myoglobin.php>.
 5. *Lipoproteins: The Medical Biochemistry Page.* Available on:
<https://themedicalbiochemistrypage.org/lipoproteins.php>.
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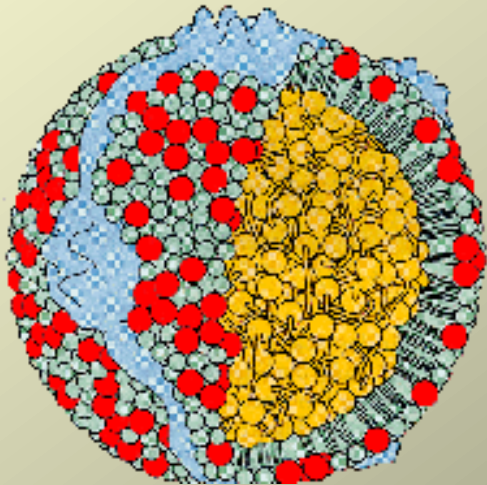
On the basis of composition, proteins are classified as **simple** or **conjugated**. **Simple** proteins contain only amino acids.

Each **conjugated** protein consists of a simple protein combined with nonprotein component.



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The nonprotein component is called a **prosthetic group**. A protein without its prosthetic group is called an **apoprotein**. A protein molecule combined with its prosthetic group is referred to as a **holoprotein**.



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CLASSIFICATION

- Glycoproteins and Proteoglycans (contain a carbohydrate component)**
 - Lipoproteins (contain lipid molecules)**
 - Chromoproteins: (contain colored component – pigment, for example hemoproteins)**
 - Metalloproteins (contain metal ions)**
 - Phosphoproteins (contain phosphate groups)**
 - Nucleoproteins (contain nucleic acids)**
- 

Hemoproteins

Oxidizable

Unoxidizable –
respiratory pigment

Fe²⁺

Hemoglobin

Myoglobin

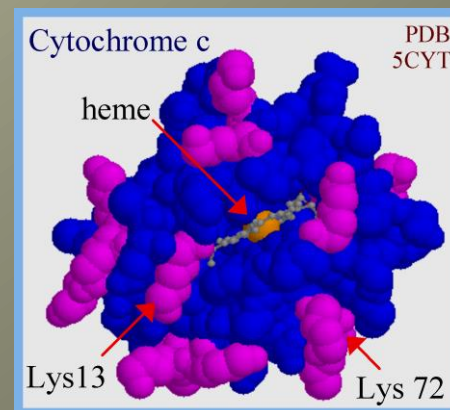
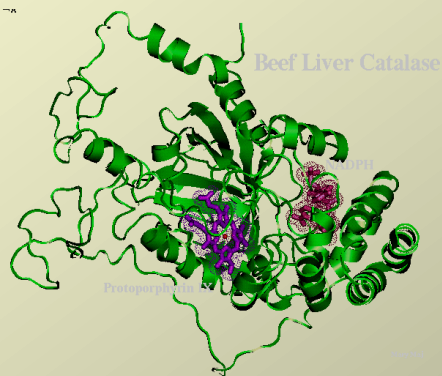
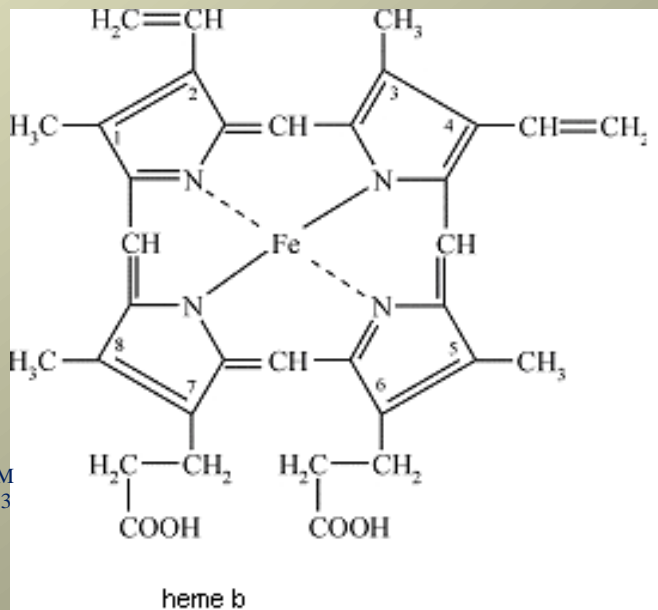
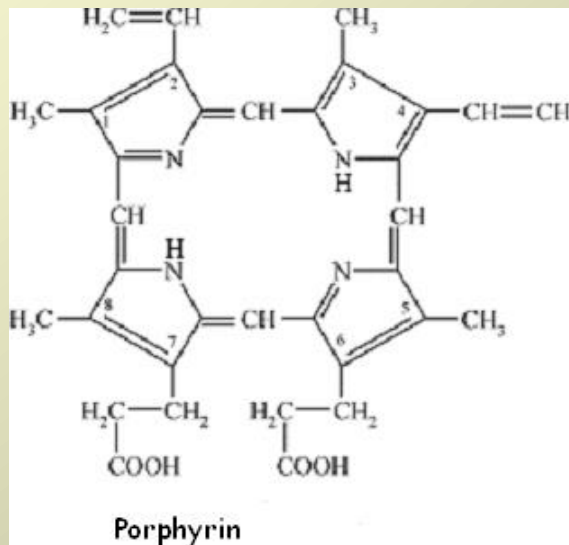
**Enzymes and
coenzymes**

Fe²⁺ ↔ Fe³⁺

Enzymes:

Catalase

**Cytochrome
and etc.**

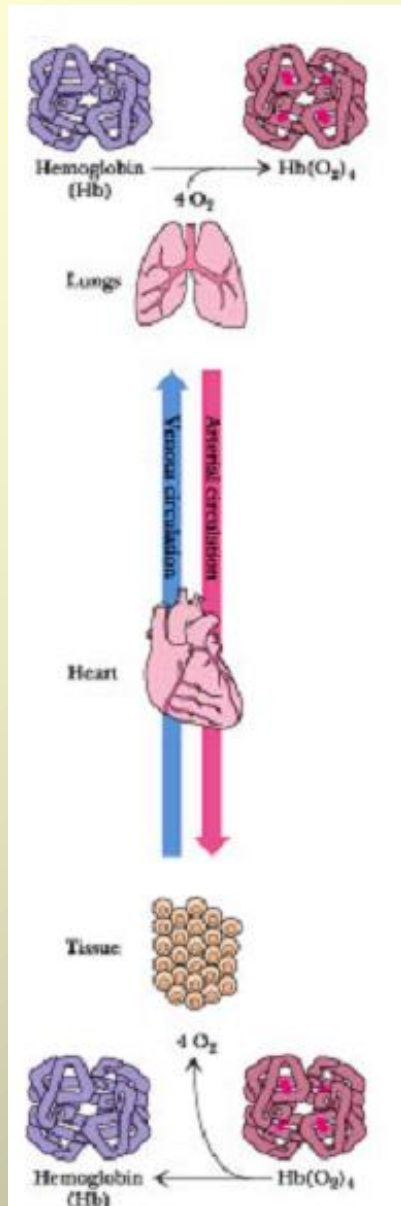


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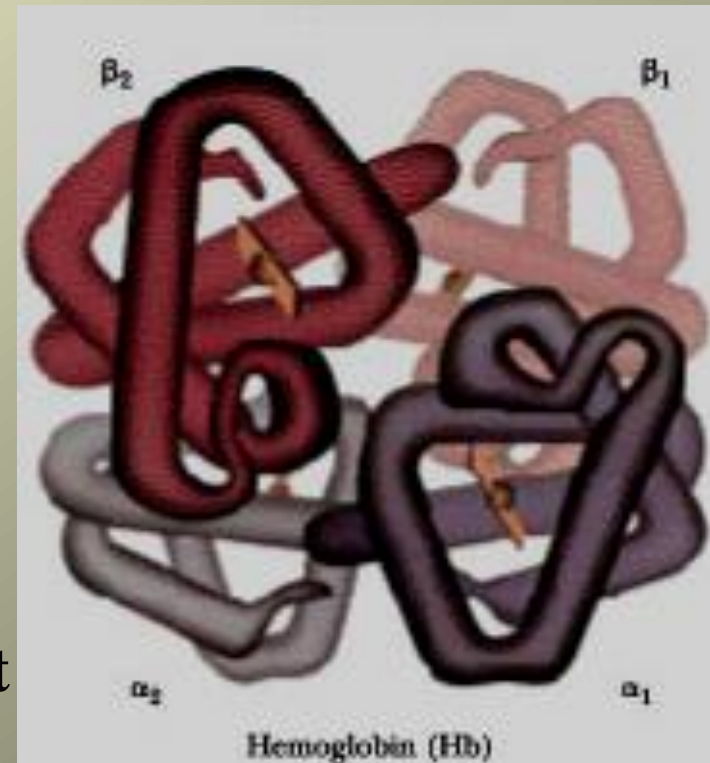
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Hemoglobin (HbA)

primary function is to transport oxygen from lungs to every tissue in the body and remove CO_2 from them. HbA is composed of two α chains and two β -chains.

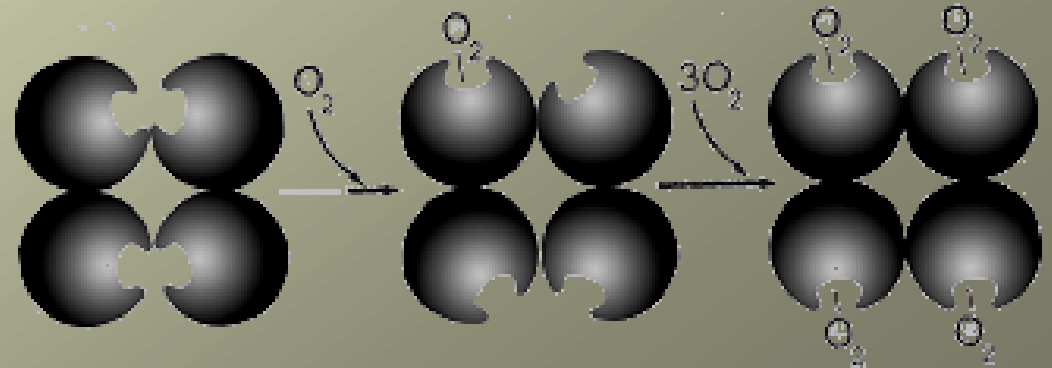
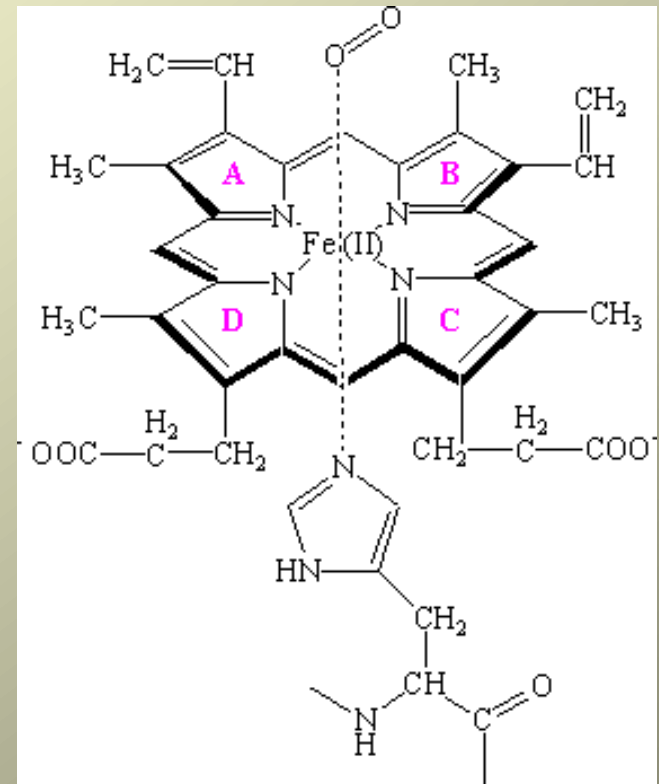
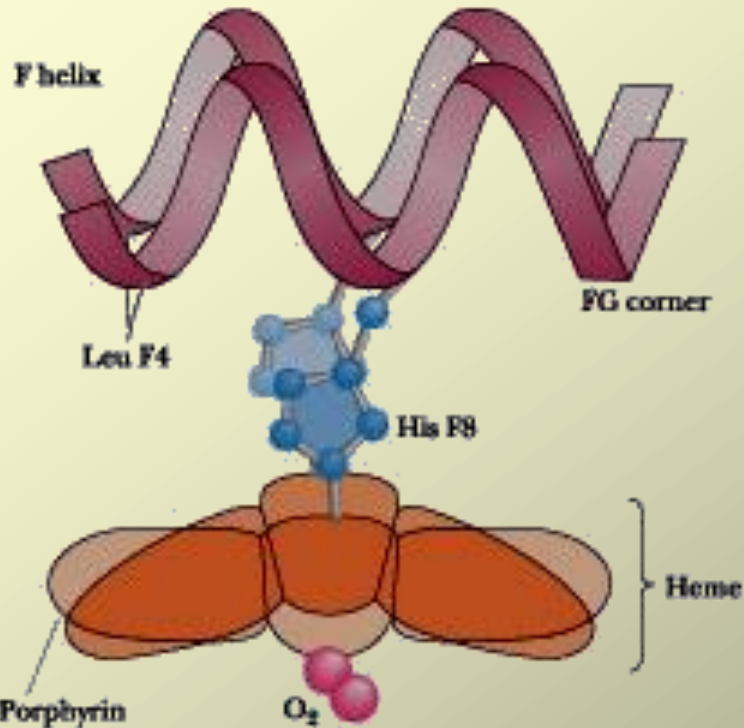


The four chains of hemoglobin, arranged in two identical subunits - $\alpha_1\beta_1$ and $\alpha_2\beta_2$ - are held together by noncovalent interactions. The association between the chains is primarily hydrophobic in nature, but hydrogen bonds and several salt bridges are also important.



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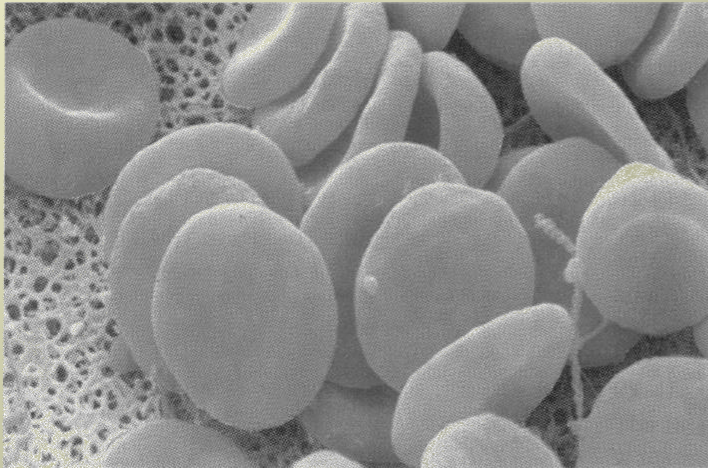
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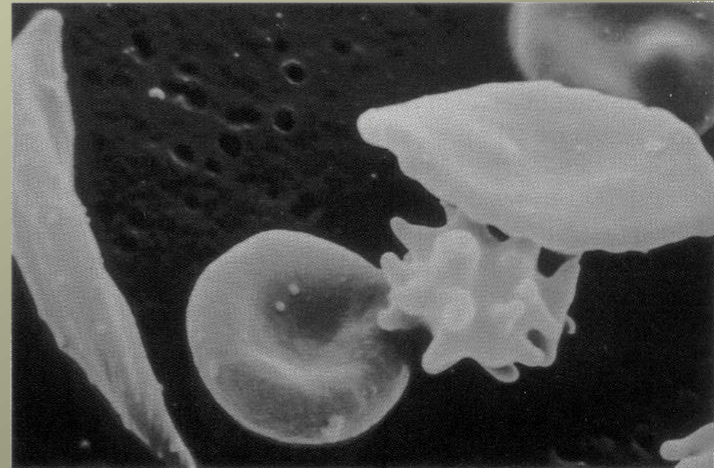
Sickle-cell anemia

Of the mutations leading to qualitative alterations in hemoglobin, the missense mutation in the β -globin gene that causes sickle cell anemia is the most common. The mutation causing sickle cell anemia is a single nucleotide substitution (A to T) in the codon for amino acid 6. The change converts a glutamic acid codon (GAG) to a valine codon (GTG). The form of hemoglobin in persons with sickle cell anemia is referred to as HbS.



(a)

2 μ m



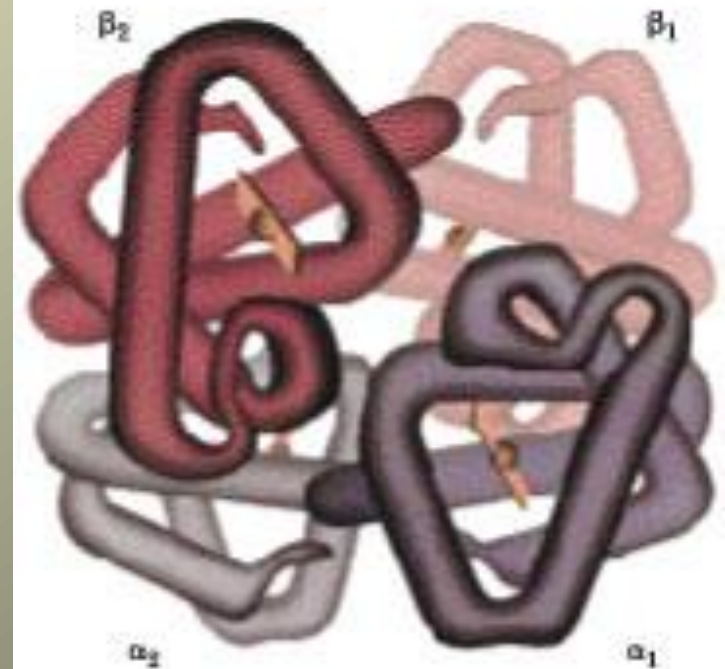
(b)

Myoglobin

Myoglobin is a monomeric heme protein found mainly in muscle tissue where it serves as an intracellular storage site for oxygen. During periods of oxygen deprivation oxymyoglobin releases its bound oxygen which is then used for metabolic purposes.



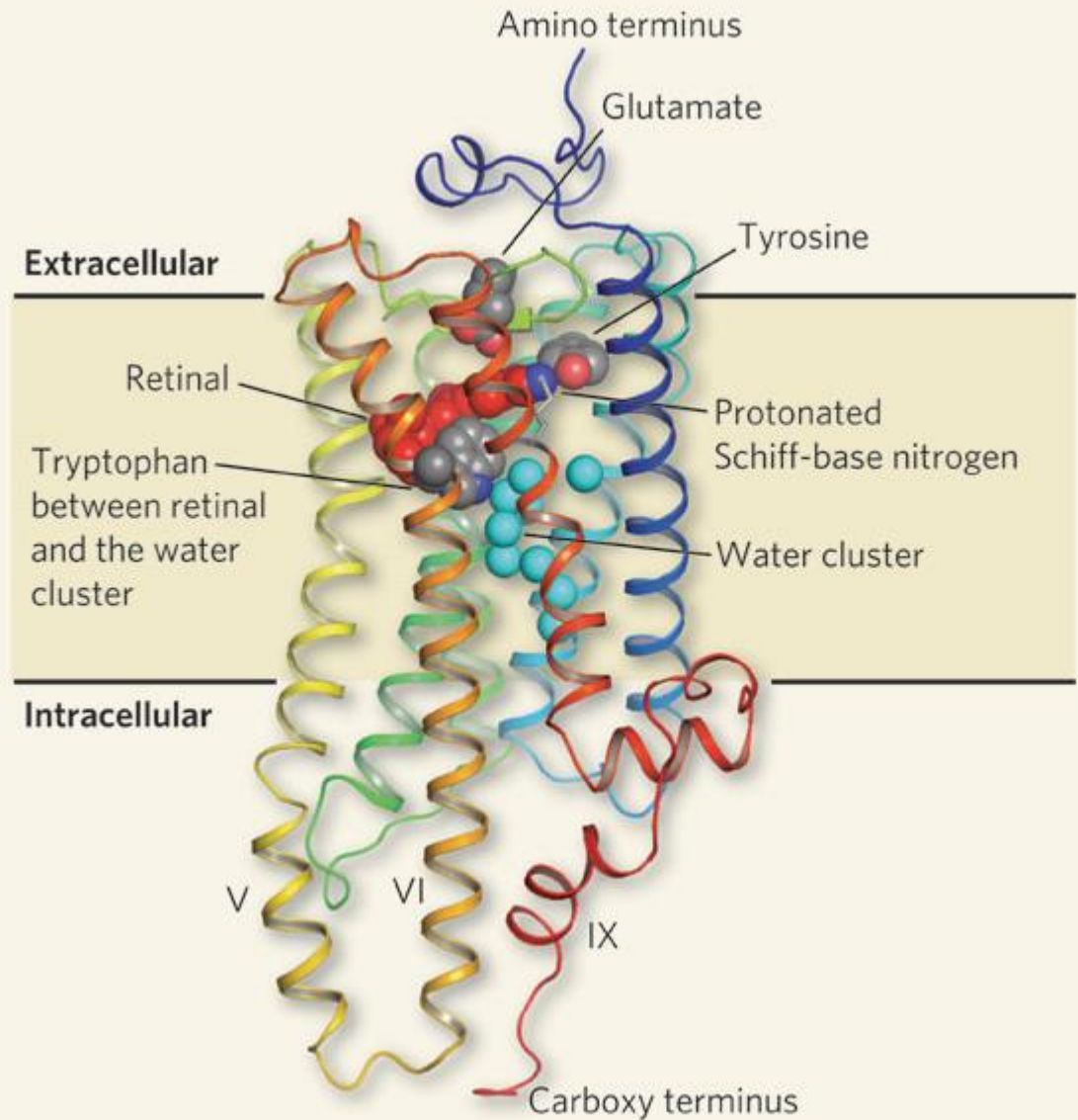
Myoglobin (Mb)



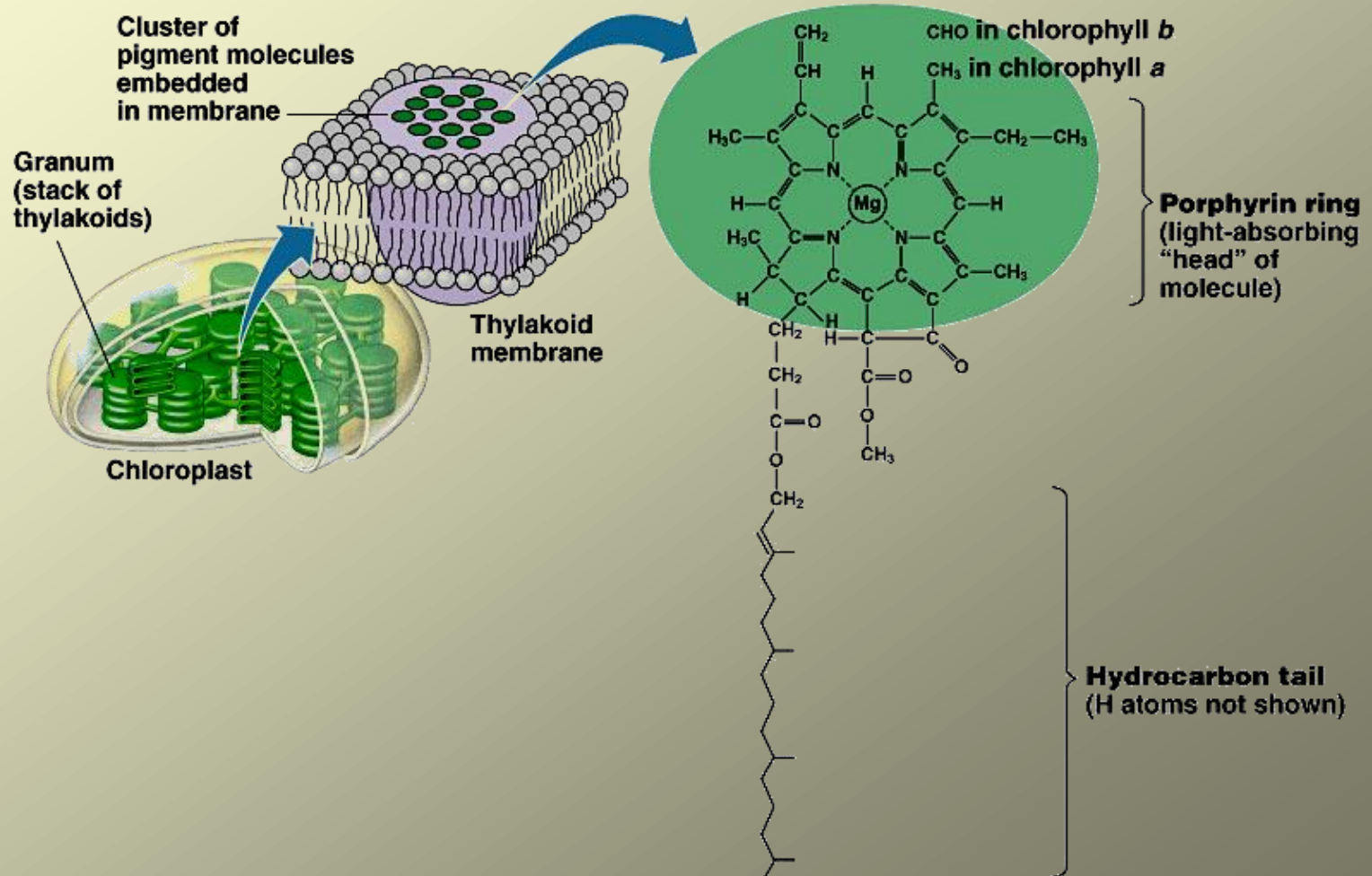
Hemoglobin (Hb)

Retinal proteins

Rhodopsin – visual pigment



Chlorophyll proteins



Cobamidproteins

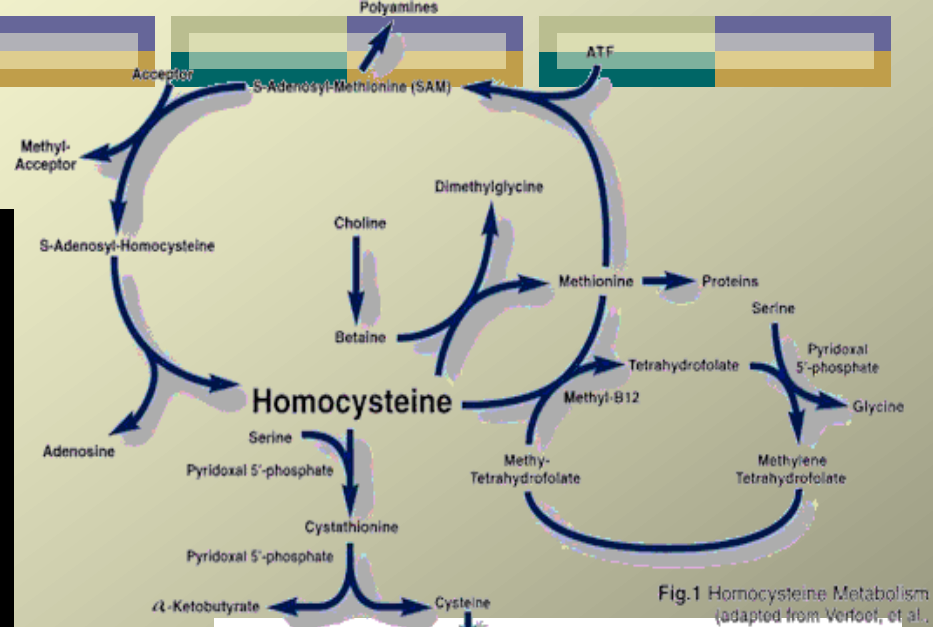
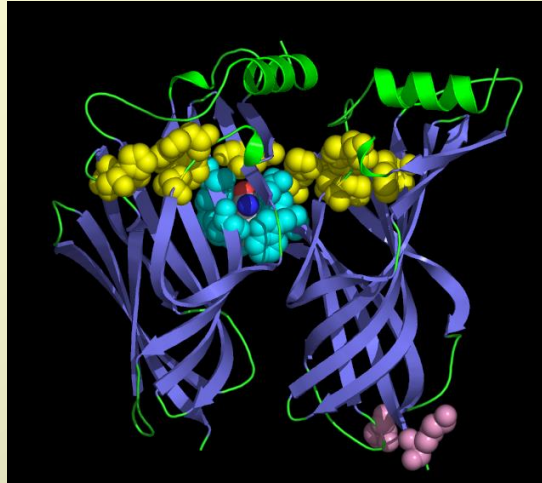
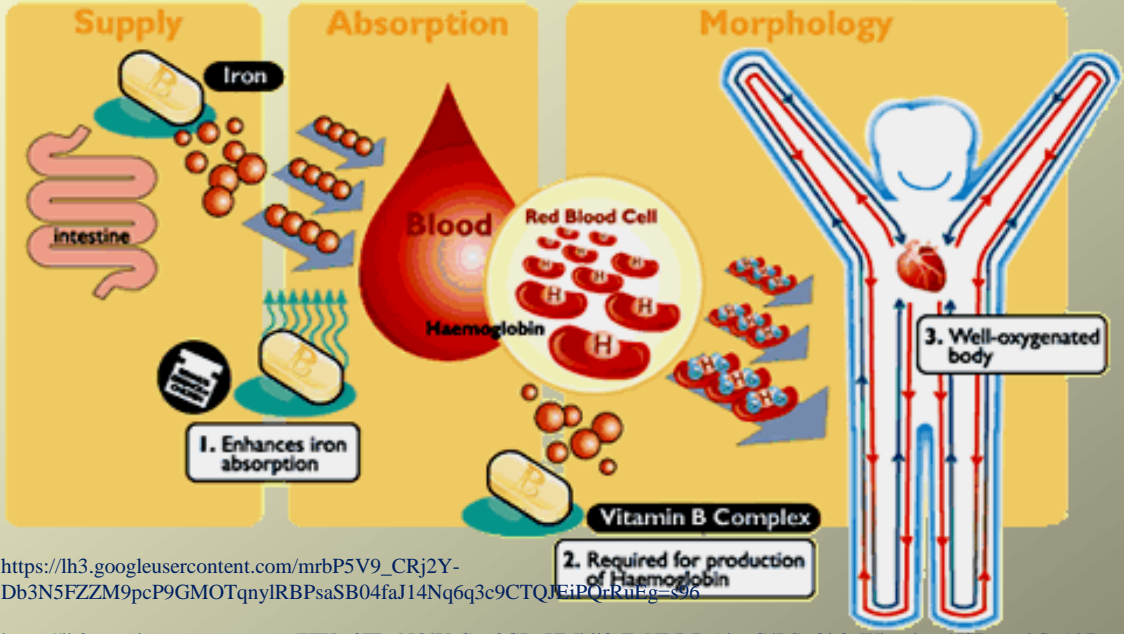
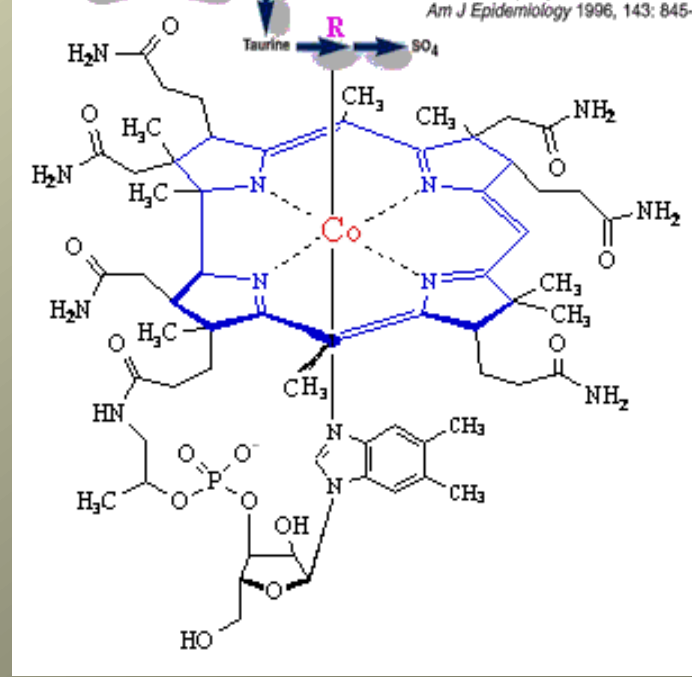


Fig.1 Homocysteine Metabolism (adapted from Verhoef, et al., Am J Epidemiology 1996, 143: 845-859)



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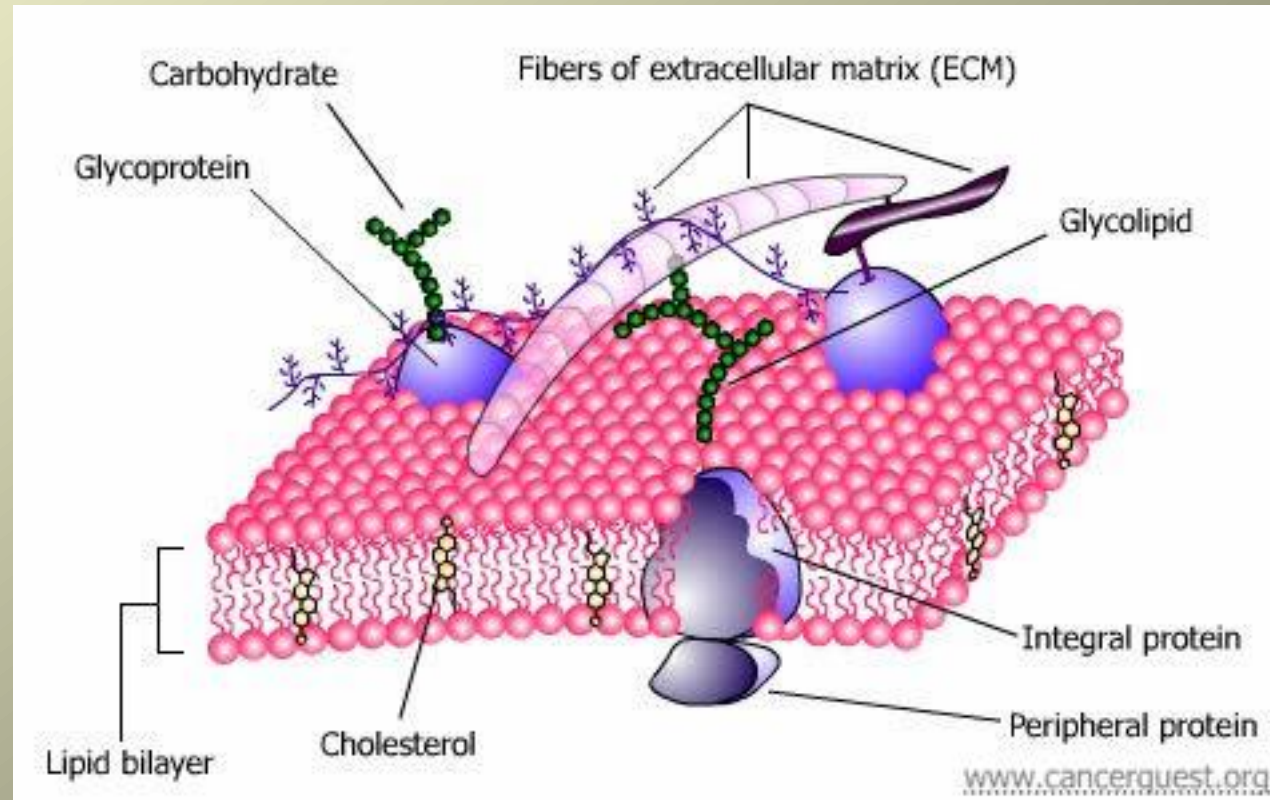
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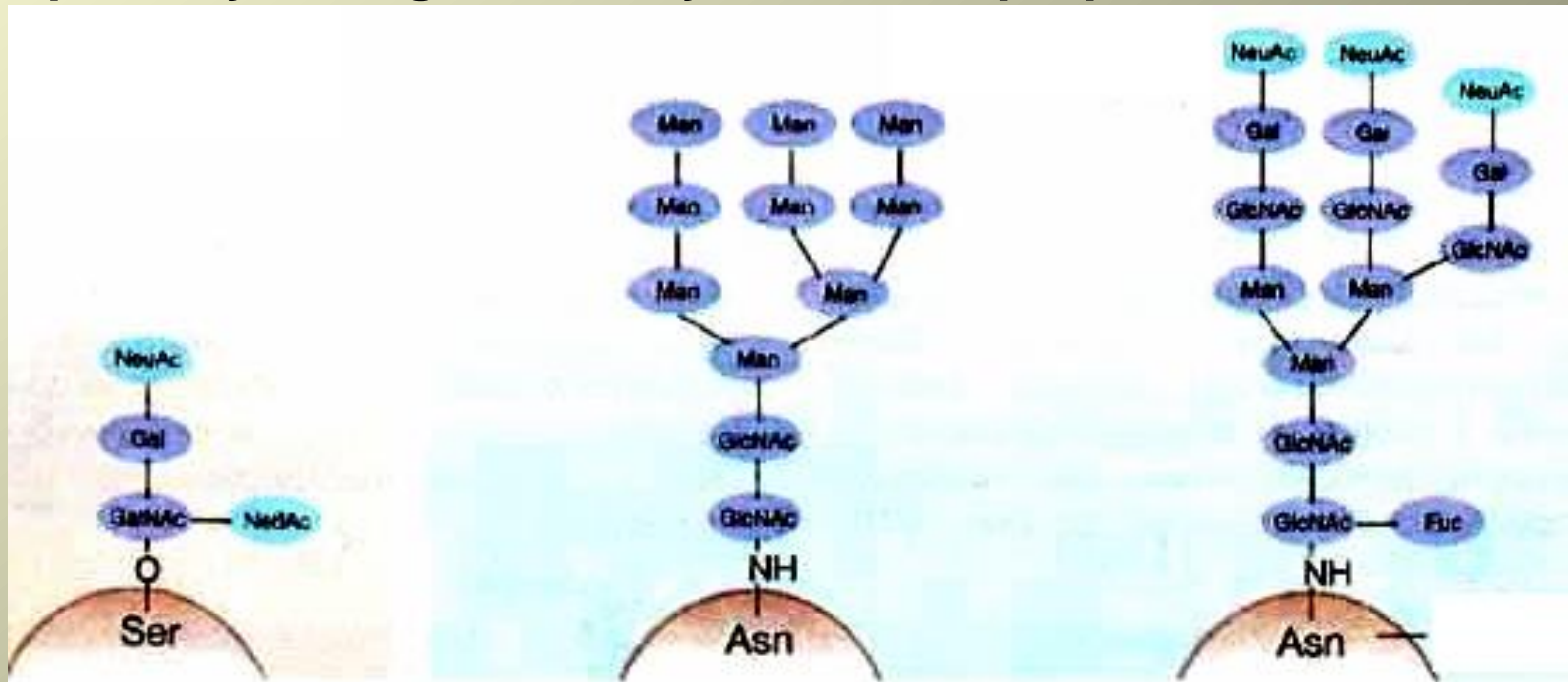
Glycoproteins

The carbohydrate chains covalently attached to glycoproteins are generally oligosaccharides of much lower molecular weight than the proteoglycans. The carbohydrate portion commonly constitutes from 1% to about 70% of a glycoprotein by weight, and never 99% as in the proteoglycans.

Glycoproteins are a diverse group of molecules that are ubiquitous constituents of most living organisms.



- Complex recognition phenomena such as cell-molecule, cell-virus, and cell-cell interactions.
- Transport proteins (transferrin, ceruloplasmin)
- Number of hormones (follicle-stimulating hormone)
- Many enzymes (ribonuclease)
- Different properties: protection from denaturation, resistance to proteolysis, high viscosity, antifreeze properties.



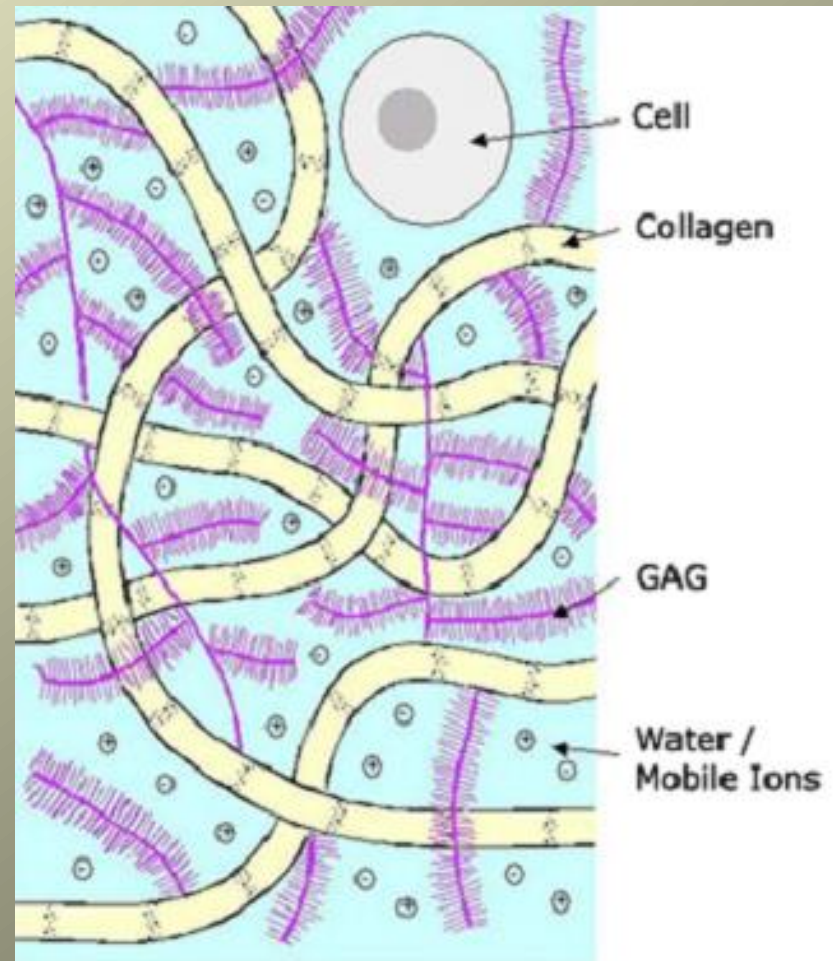
Proteoglycans

This molecules are found predominantly in the extracellular matrix of tissues.

All proteoglycans contain **glycosaminoglycans (GAGs)**.

The specific GAGs of physiological significance are:

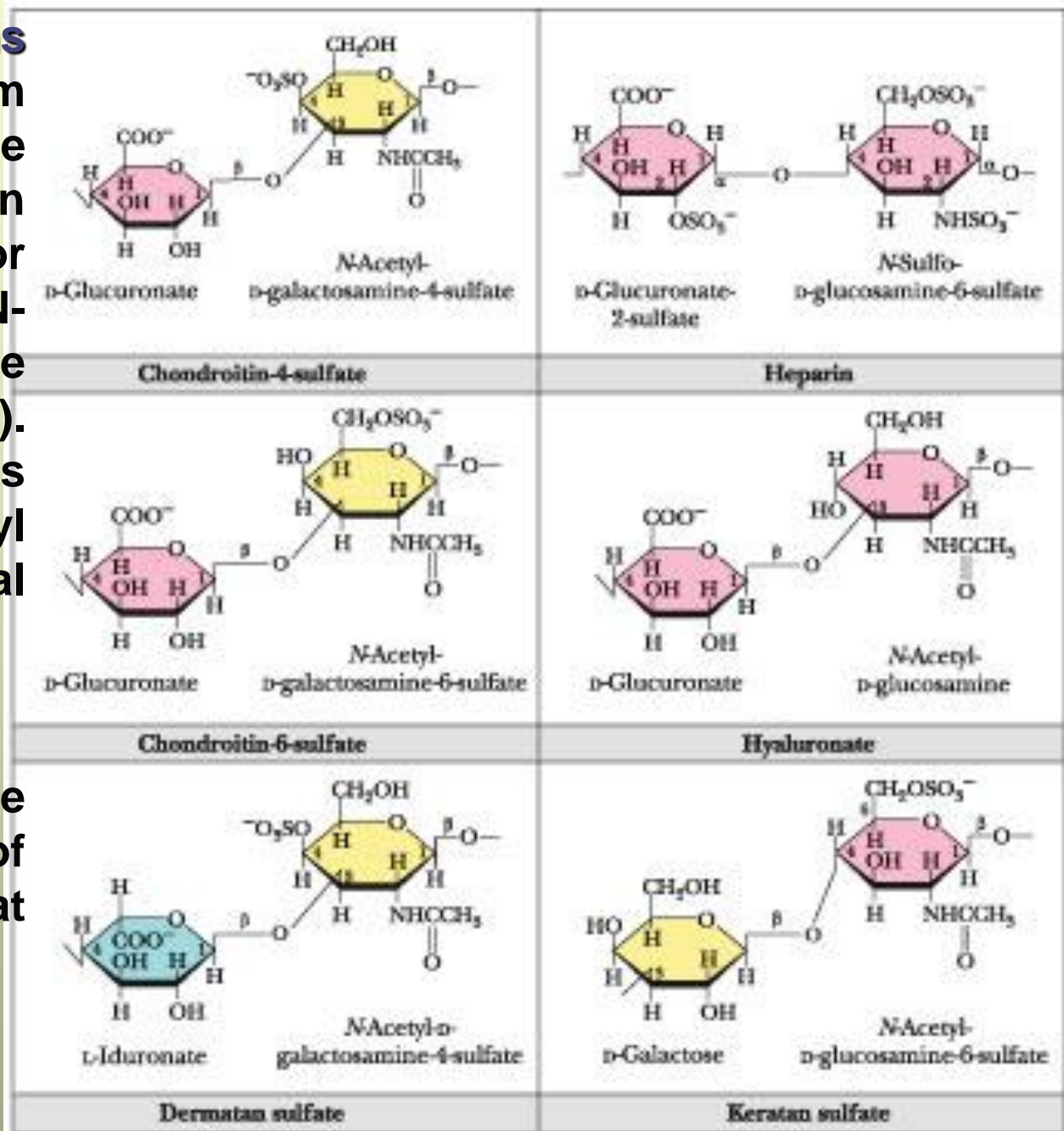
hyaluronic acid,
dermatan sulfate,
chondroitin sulfate,
heparin, heparan sulfate,
and **keratan sulfate.**



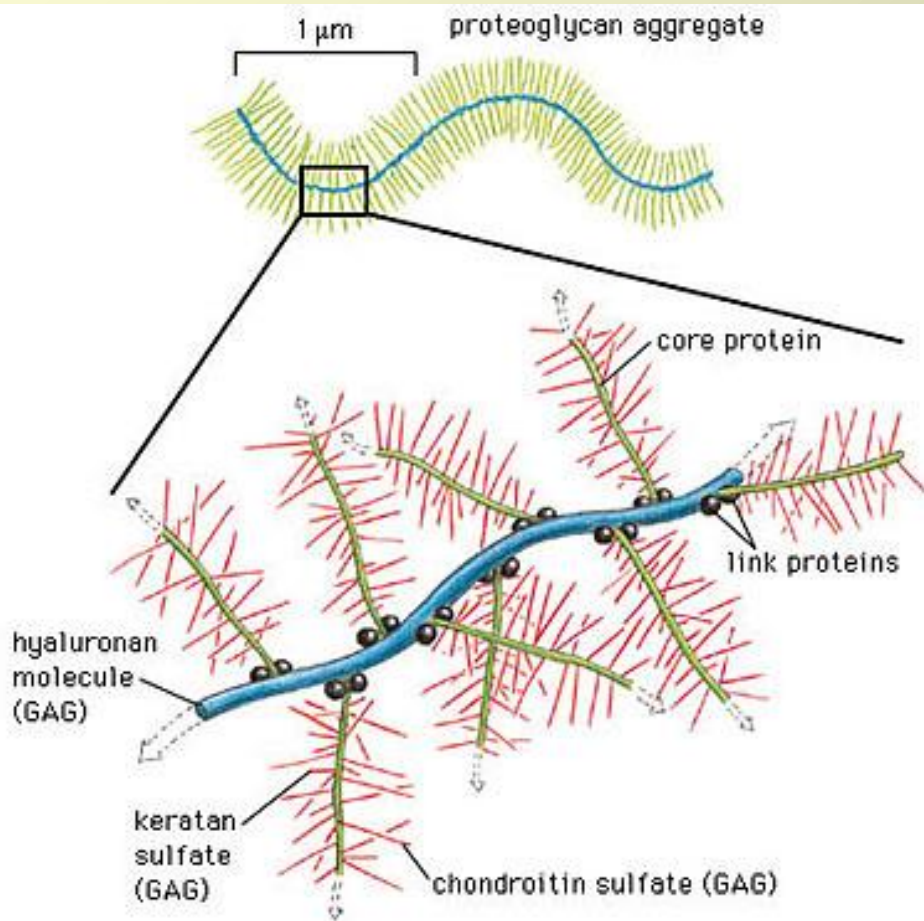
Glycosaminoglycans

are formed from repeating disaccharide arrays. This units contain a hexuronic acid (or uronic acid) and N-acetylhexosamine sulfate (or N-acetylglucosamine). Many disaccharide units contain both carboxyl and sulfate functional groups.

All the GAGs therefore have large numbers of negative charges at physiological pH.



Along with the high viscosity of GAGs comes low compressibility, which makes these molecules ideal for a lubricating fluid in the joints. At the same time, their rigidity provides structural integrity to cells and provides passageways between cells, allowing for cell migration.



Hyaluronates are important components of the vitreous humor in the eye and of synovial fluid, the lubricant fluid of joints in the body.

The chondroitins and keratan sulfate are found in tendons, cartilage, and other connective tissue, whereas dermatan sulfate, as its name implies, is a component of the extracellular matrix of skin.

Heparin, is a natural anticoagulant substance. It binds strongly to antithrombin III and inhibits blood clotting.

Kristle domain

GlcNAc

IdoA6S

GlcNAc

IdoA6S

IdoA6S

N-domain

The Effect of Heparin on the Blood Vessels

Initial Condition

Large amounts of clotting agents move through the bloodstream. Blood begins to leak through a small defect in the vessel wall.



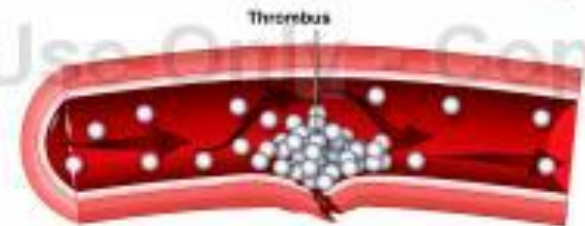
Effective Clotting

Clotting agents collect at the defect site, effectively stopping the leakage of blood.



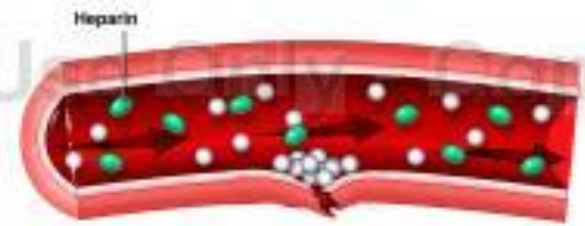
Thrombus Formation

When too many clotting agents accumulate, a thrombus forms, partially blocking blood flow.



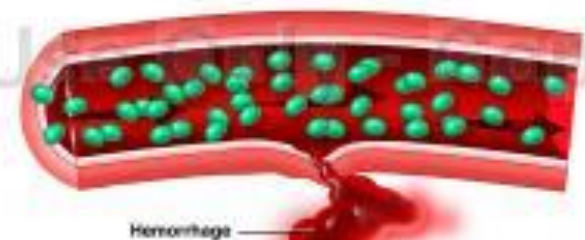
Low Dose Heparin

Low doses of heparin clear the thrombus yet leave enough of the clotting agents for the effective closure of the vessel wall defect.



High Dose Heparin

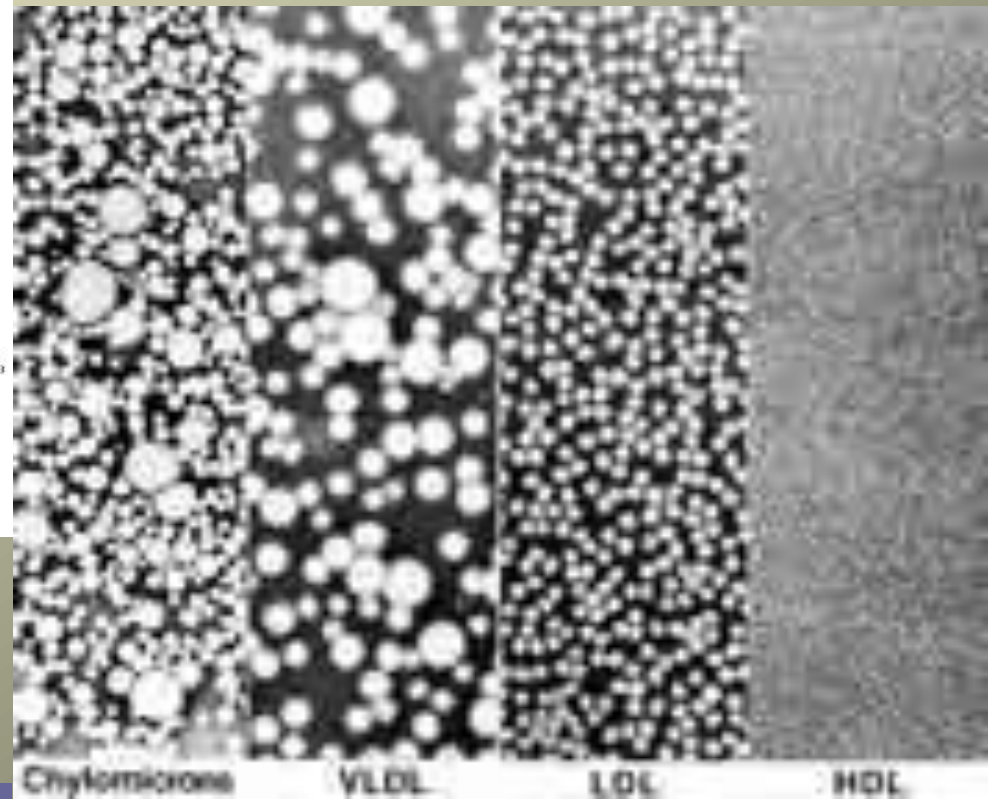
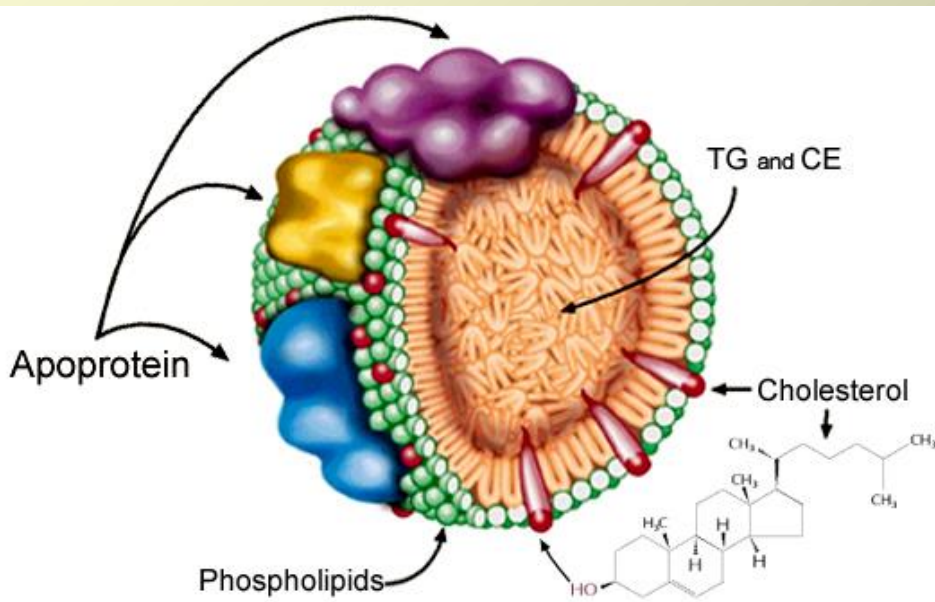
High doses of heparin completely eliminate clotting agents. This leads to uncontrolled hemorrhage through the vessel wall defect.



LIPOPROTEINS

group of molecular complexes found in the blood plasma of mammals.

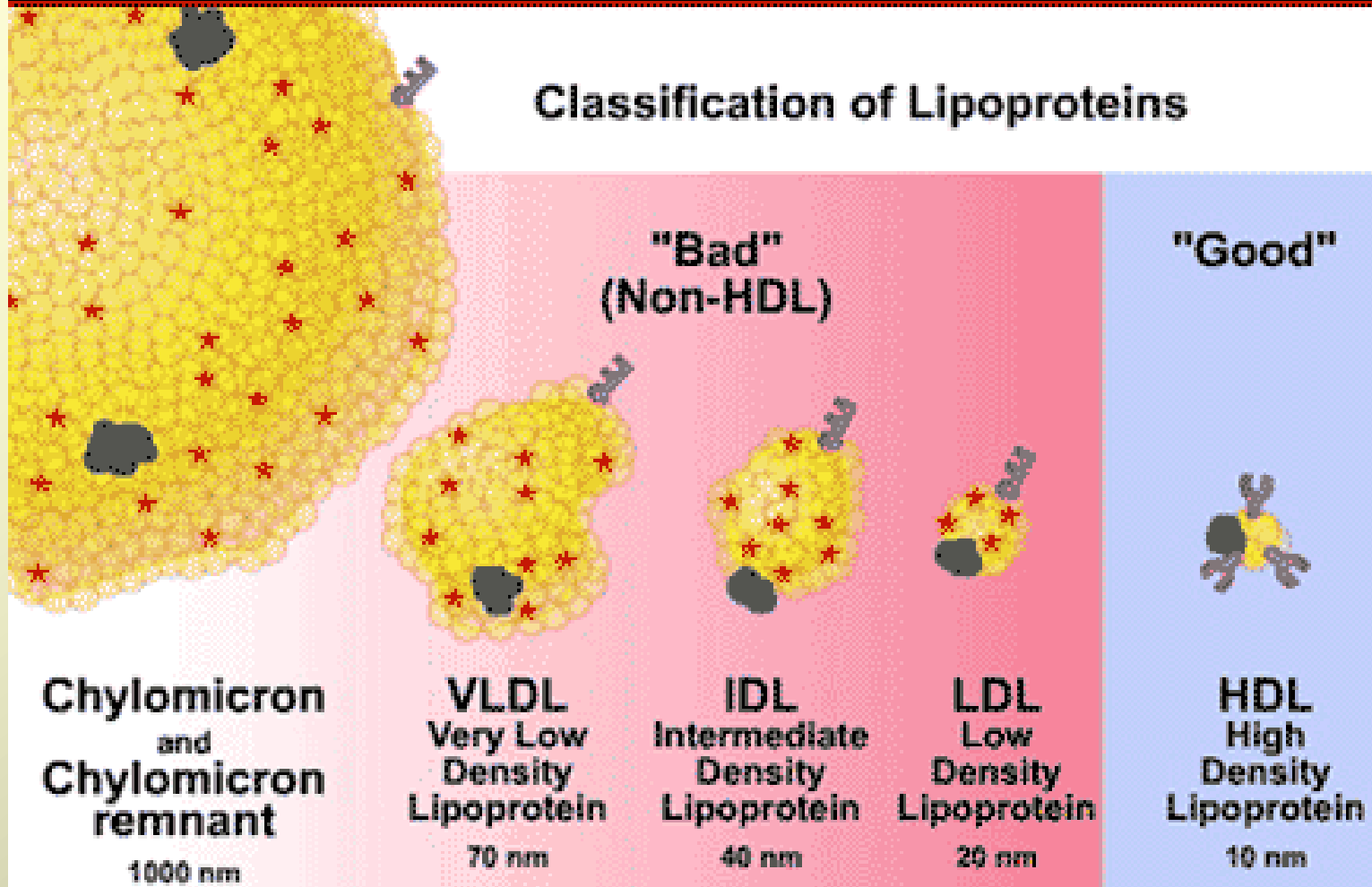
Plasma lipoproteins transport lipid molecules (triacylglycerols, phospholipids, and cholesterol) through the bloodstream from one organ to another.



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Classification of Lipoproteins



Lipoproteins differ in the ratio of protein to lipids, & in the particular apoproteins & lipids that they contain.

They are classified based on their density:

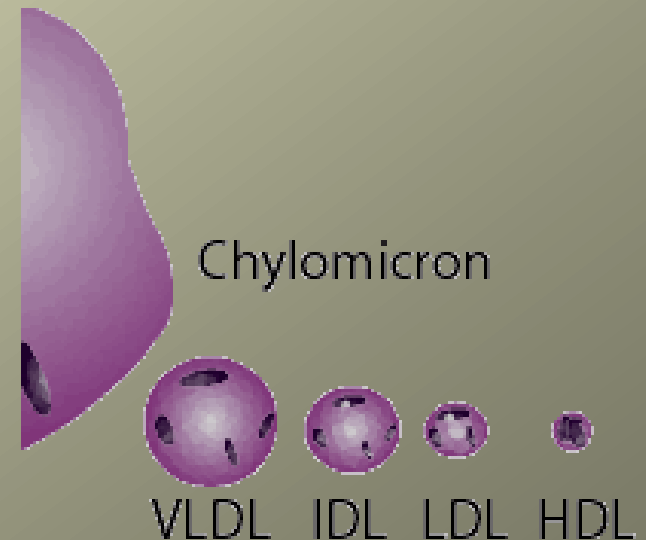
Chylomicron (largest; lowest in density due to high lipid/protein ratio; highest % weight triacylglycerols)

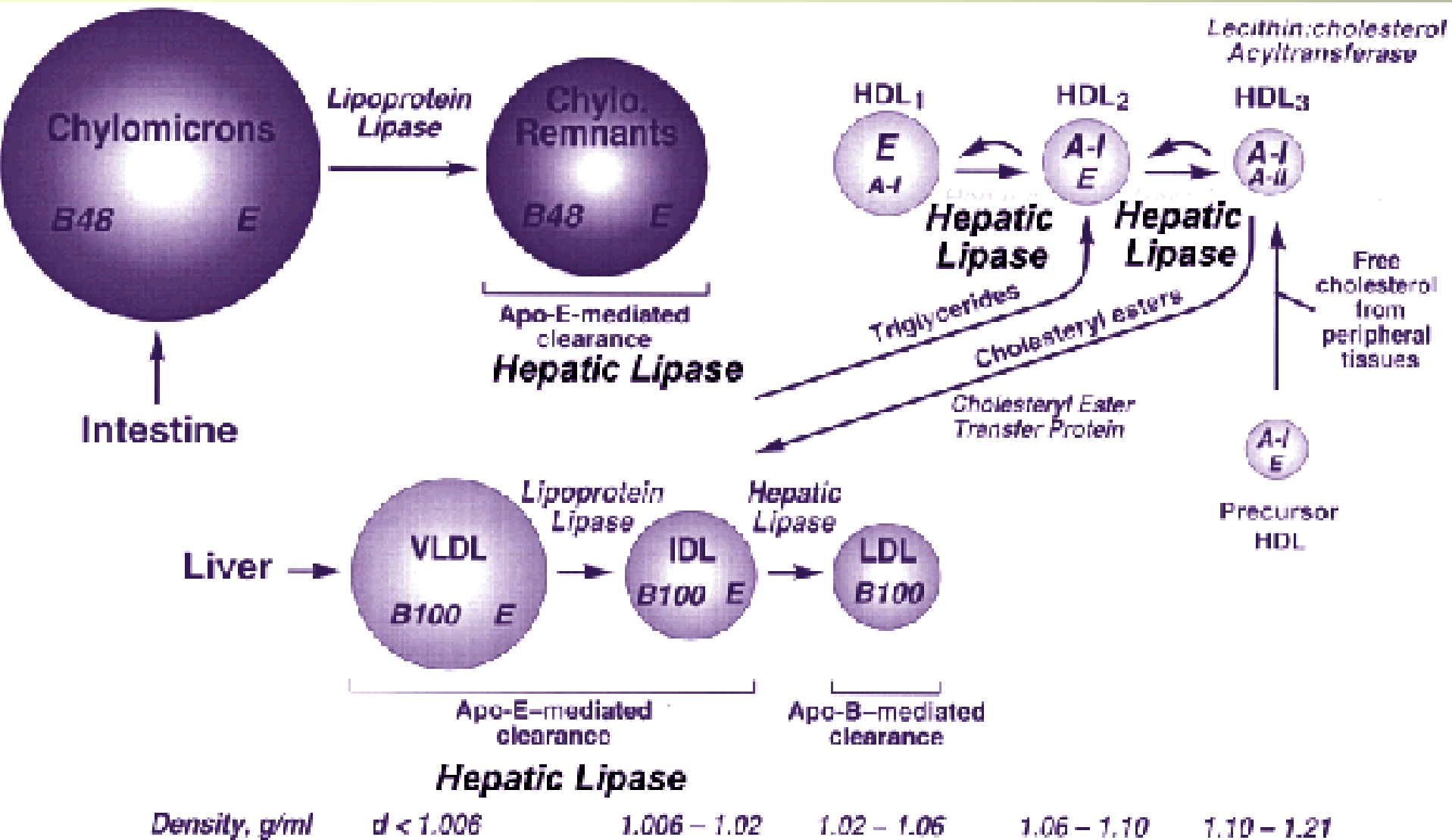
VLDL (very low density lipoprotein; 2nd highest in triacylglycerols as % of weight)

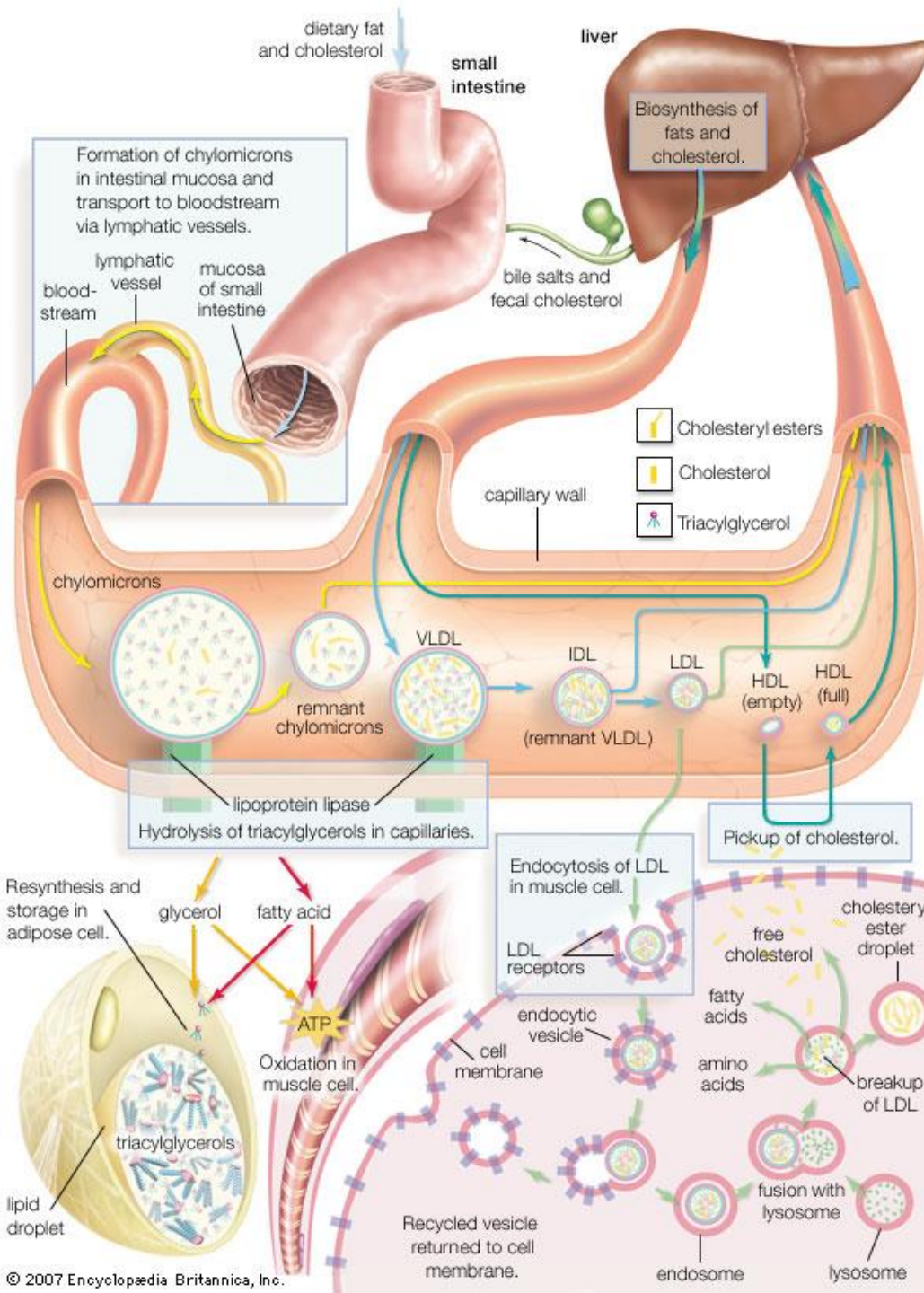
IDL (intermediate density lipoprotein)

LDL (low density lipoprotein, highest in cholesteryl esters as % of weight)

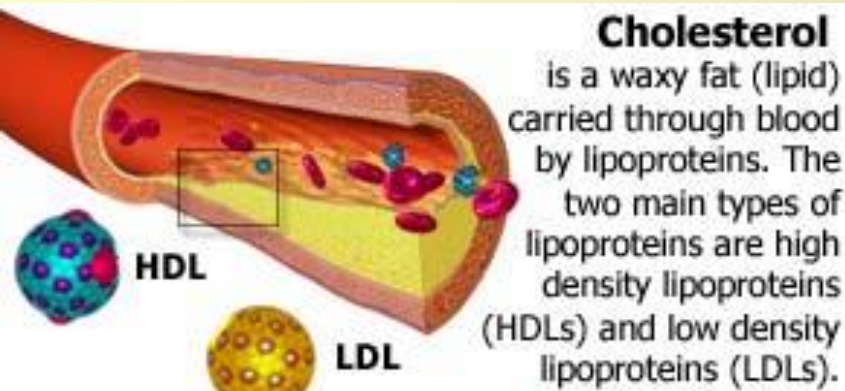
HDL (high density lipoprotein; highest in density due to high protein/lipid ratio)



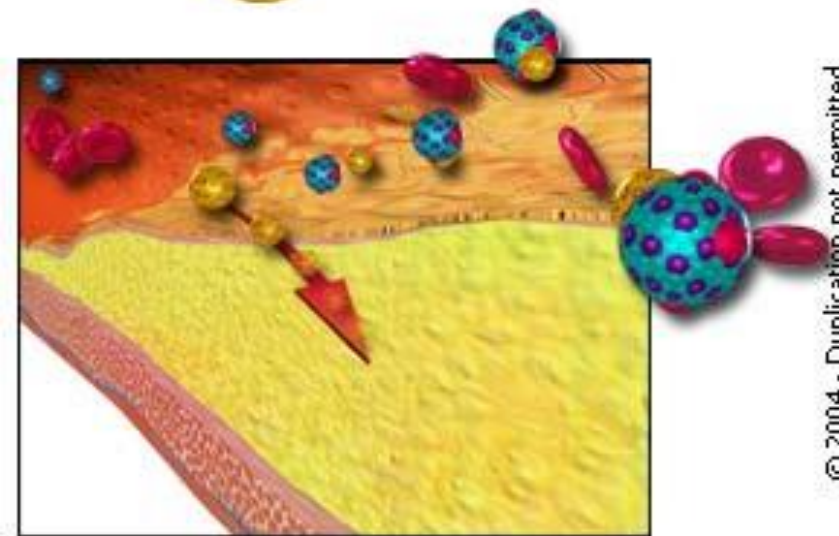




Lipoproteins and atherosclerosis



LDL и VLDL – atherogenic
HDL- antiatherogenic



Normal coronary artery



Atherosclerosis



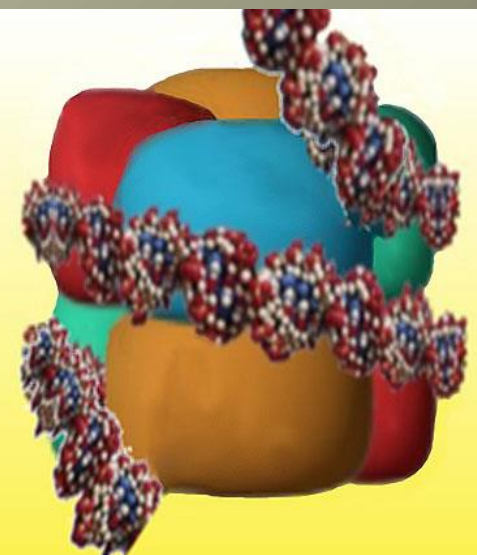
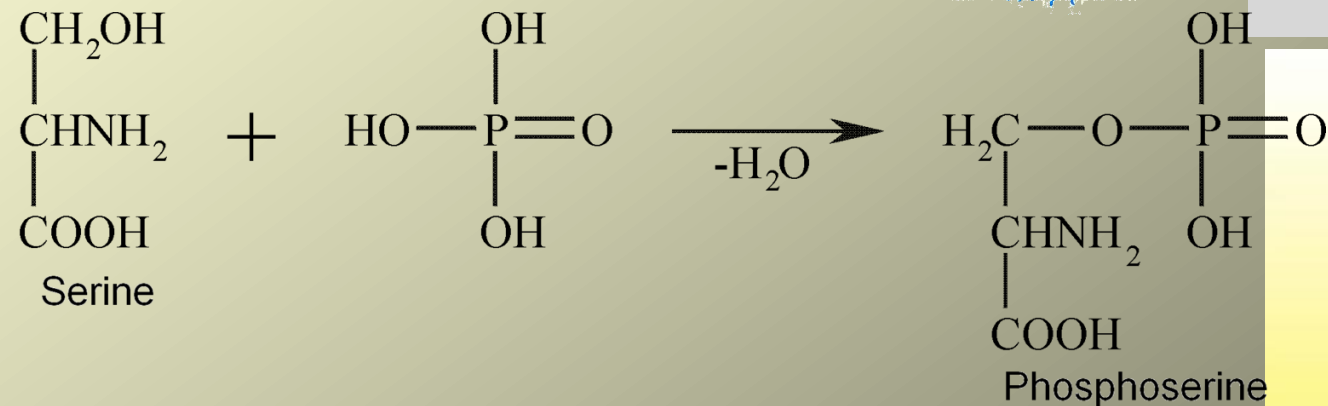
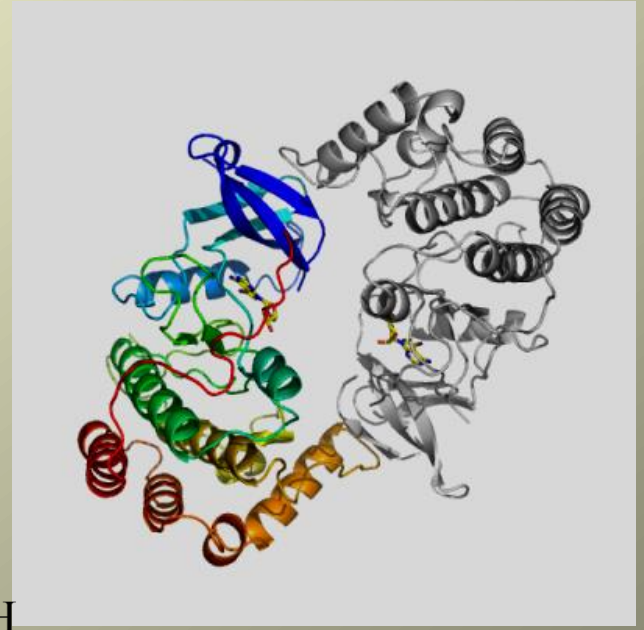
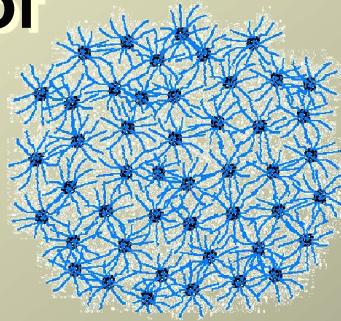
Atherosclerosis with blood clot



HDLs (good cholesterol) carry LDLs (bad cholesterol) away from artery walls. LDLs stick to artery walls and can lead to plaque build-up (atherosclerosis).

Phosphoproteins

- 1) Nutritious (casein)
- 2) Structural (histones)
- 3) Enzymes (regulation of the enzyme activity)



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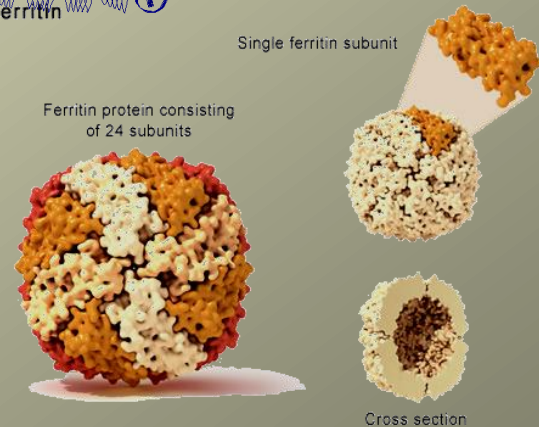
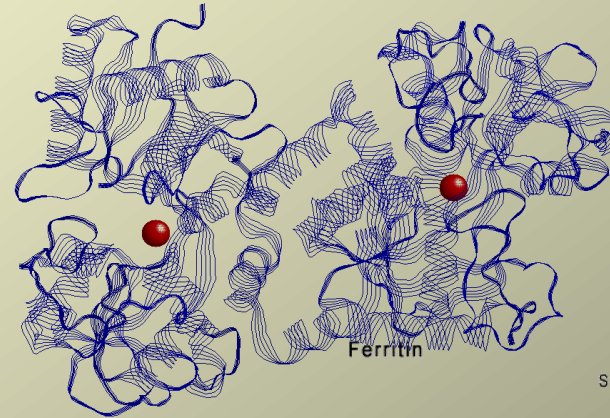
Metalloproteins

1) Transport (transferrin)

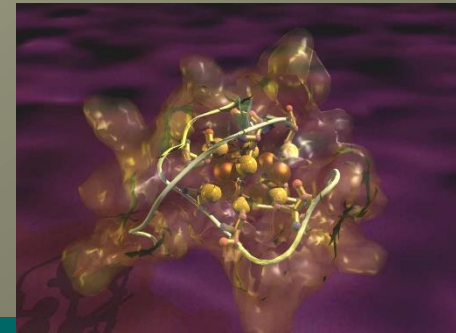
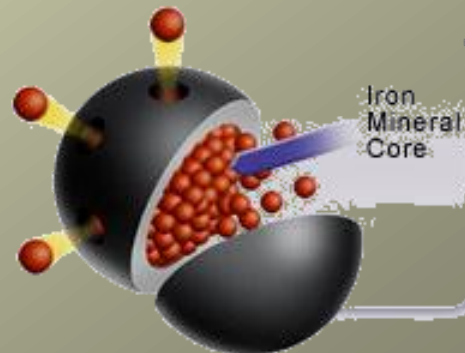
2) Storage (ferritin)

3) Enzymes (alcohol dehydrogenase)

4) Neutralizing



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Conclusions

1. Based upon the chemical composition proteins are divided into two major classes: simple proteins, which are composed of amino acids and additional organic and inorganic groupings.
 2. Conjugated proteins include glycoproteins, which contain carbohydrates, lipoproteins, which contain lipids, chromoproteins, which contains pigments, etc.
 3. In living organisms conjugated proteins serve a great amount of functions.
- 

Do you have any questions?

Thank you for your attention!

