

Major types of nutrients and digestion

Lecture 1+2

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- The 7 types of nutrients are
① Carbohydrates, ② Protein, ③ Fats, Vitamins,
Minerals, Water and Fibers

the
major

every type of food
has different serving.

We should have every
day

examples of
serving size:

* Grain products—1
slice of white bread
or 1/2 cup of
cooked rice;

* Vegetable
group—1/2 cup
cooked vegetables

* Fruit group—1
apple or banana;

* Milk Group—1
cup of milk or 60g
processed cheese;

* Meat and Beans
Group—2-3 oz (60-
80g) cooked lean
meat or fish or 1
egg or 2 tbsp
peanut butter.

table
spoon

Food Guide Pyramid

A Guide to Daily Food choices

Fats, Oils, & Sweets
USE SPARINGLY

as much as less

KEY
□ Fat (naturally occurring and added) ■ Sugars (added)

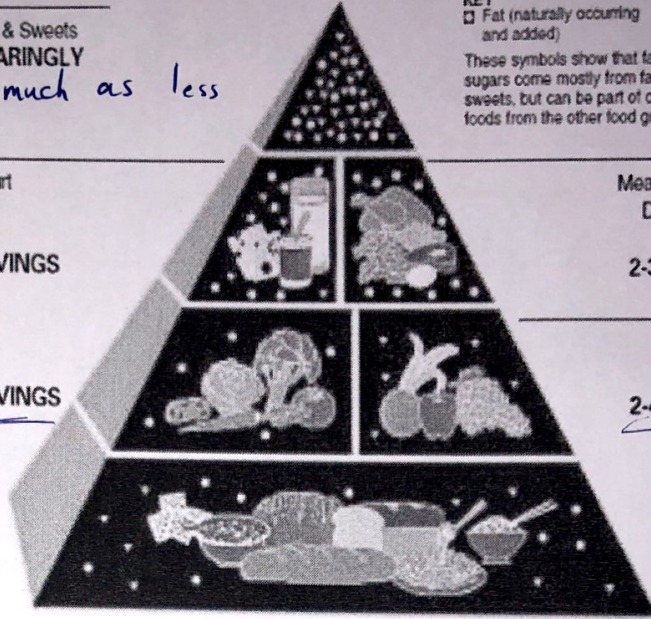
These symbols show that fat and added sugars come mostly from fats, oils, and sweets, but can be part of or added to foods from the other food groups as well.

Milk, Yogurt
& Cheese
Group
2-3 SERVINGS

Meat, Poultry, Fish,
Dry Beans, Eggs
& Nuts Group
2-3 SERVINGS

Vegetable
Group
3-5 SERVINGS

Fruit
Group
2-4 SERVINGS



1. Carbohydrates

Functions

- Energy
- Fibers *undigested carbs*
- Recognition and adhesion between cells

Carbohydrate covalently attached to proteins or lipids examples: glycoproteins, glycolipids, these classes of molecules are called glycoconjugates.

bound
to lipid

bound
to proteins

- Most foods derived from animals, such as meat or fish, contain very little carbohydrate except for small amounts of glycogen for example Liver contains 5% carbohydrates
- The major dietary carbohydrate of animal origin is lactose.
- Although all cells require glucose for metabolic functions, neither glucose nor other sugars are specifically required in the diet.

Oxidation

- 1g carbohydrates produces about 4 kcal/g,
- 1g proteins produces 4 kcal/g
- 1g fats produces 9 kcal/g. → this is why it's the major thing for energy
- Energy is also expressed in joules. One kilocalorie equals 4.18 kilojoules (kJ).

له الجسم يمتزج السكر على شكل glycogen في الكبد، (العظام) لكن يخزن بدائى عند (starvation) ويبدا سرقا الدهون

Carbohydrates are classified into three groups:

1-Monosaccharide's: (simplest)

- = Also called simple sugars
- = Have the formula $(CH_2O)_n$
- = Cannot be broken down into smaller sugars.

2-Oligosaccharides:

- = Consist of from **2 to 10 monosaccharides** molecules joined by a linkage called glycosidic bonds (covalent bond).
- = The most abundant are the disaccharides
- = Trisaccharides also occur frequently.
- = All common monosaccharides and disaccharides have names ending with the suffix "-ose."

3- Polysaccharides: are polymers of monosaccharide's.

They may be either linear like cellulose or branched like glycogen. Polysaccharides may contain hundreds or even thousands of monosaccharide units.

more than
10

2 monosaccharides

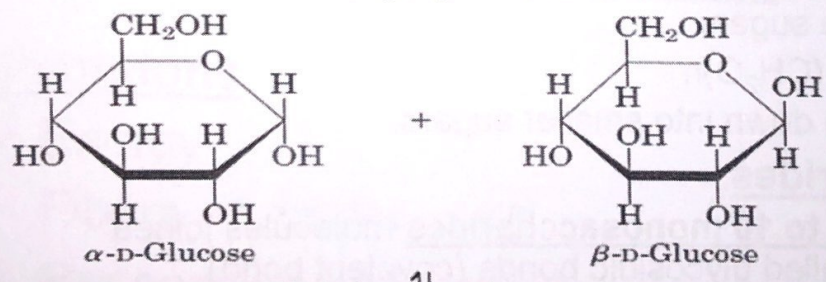
Disaccharides

the building block of it is glucose but different in site of bond

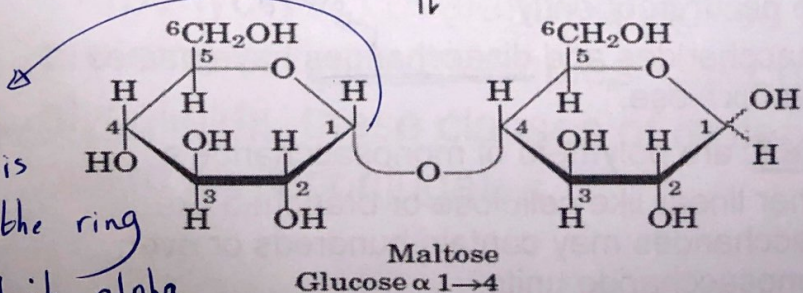
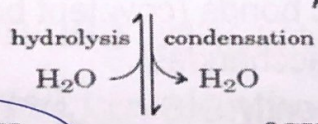
- 1- **Maltose** Consist of two glucose units joined by α -1,4. Carbon number 1 + C4 of next glucose
Comes from the hydrolysis of starch
- 2- **Cellobiose**, is a two glucose disaccharide joined by β -1,4 obtained from the acid hydrolysis of cellulose
- 3- **Isomaltose** consists of two glucose units linked by α -1,6 obtained from the hydrolysis of some polysaccharides like dextran
- 4- **Trehalose** two glucose units linked by an α -bond (α -1,1)
- 5- **Lactose** Milk sugar, Composed of Galactose & Glucose joined by (β -1,4) link.
- 6- **Sucrose** Composed of (glucose- α -1,2-fructose) obtained from cane or beet commonly known as table sugar

Glycosidic bond in maltose

Glycosidic bond the covalent bond between the anomeric carbon atom of a saccharide and some other group or molecule with which it forms a glycoside.



alpha 1,4 glycosidic bonds are formed when the OH on the carbon-1 is below the glucose ring while beta 1,4 glycosidic bonds are formed when the OH is above the plane



If the bond is down the ring we call it alpha
If it above the plane of ring called Beta.

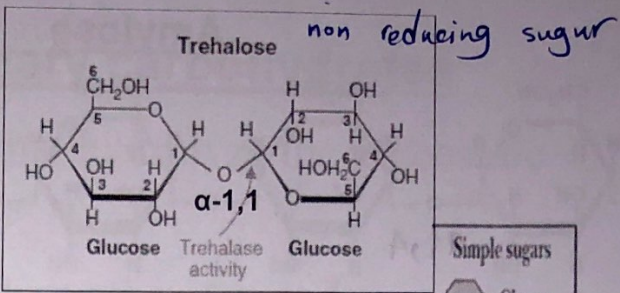
Carbohydrates as monosaccharides

- Aldoses \rightarrow Aldehyde (functional) group in C1
- Ketoses \rightarrow Ketone in C2

carbon

Reducing and non-reducing end
non bonded

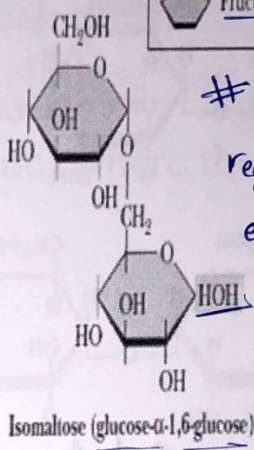
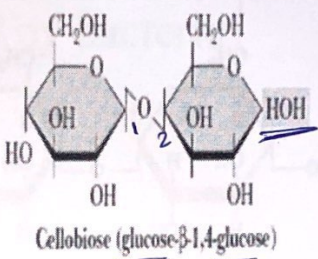
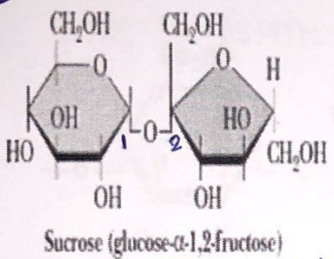
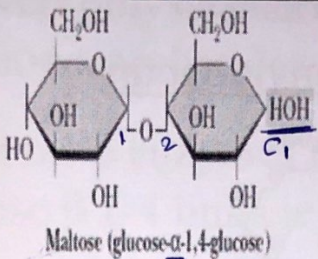
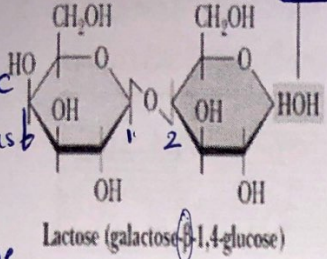
Free anomeric carbon (reducing end)



Simple sugars

- Glucose
- Galactose
- Fructose

glycosidic bond must have at least one anomeric carbon



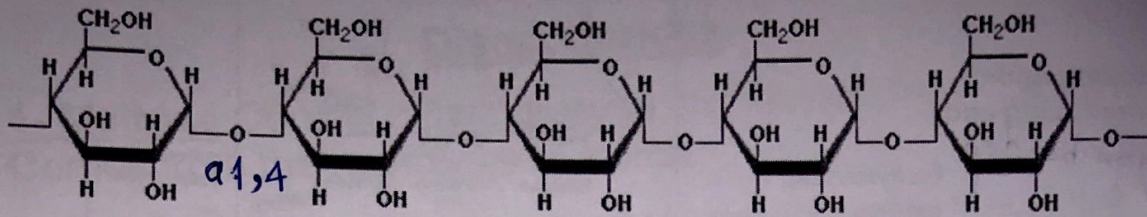
Every bond required special enzyme to break
reducing

Aldoses \rightarrow Ketoses
so non reducing sugar

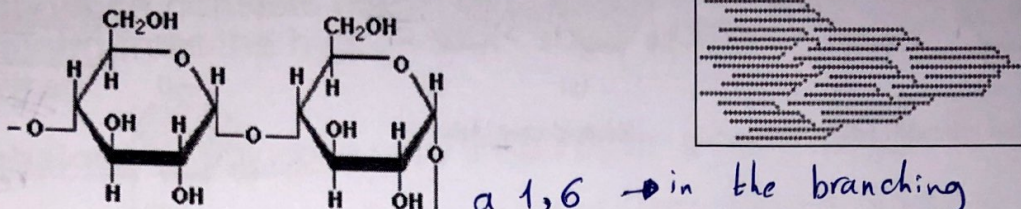
Polysaccharides

- **Starch** Starch represents the main type of digestible polysaccharides. Found in cereal grains (wheat, rice, corn, oats, and barley) and tubers such as potatoes.
- **Starch is made up of :-** *glucose mainly*
 - 1- **Amylose** (about 20% of starch) strait chain of glucose which is linked by α -1,4 glycosidic bonds
 - 2- **Amylopectin** (about 80% of starch) (branched) the α -1,4 chains contain branches connected via α -1,6 glycosidic bonds
- **None starch polysaccharides are the main component of dietary fibers which include:** Cellulose, hemicellulose, pectin's and gum.

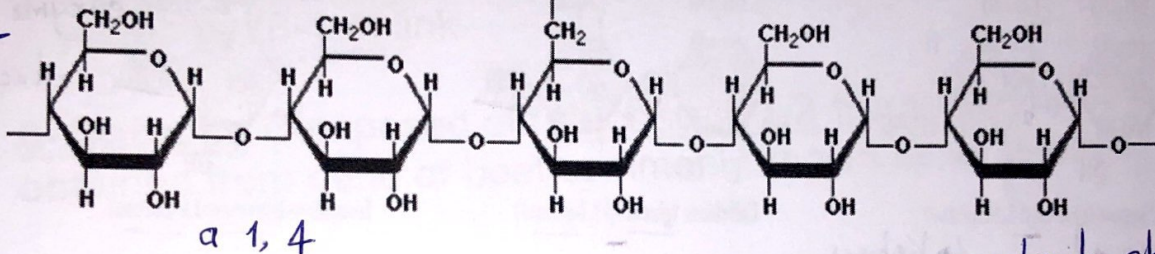
Amylose



Amylopectin



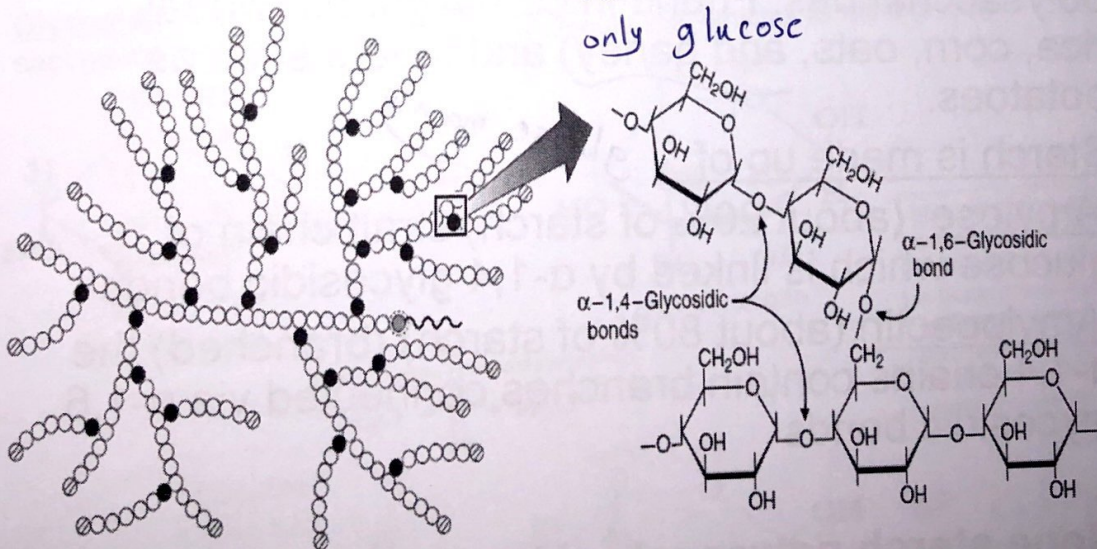
branched one



- both Amylose + Amylopectin form starch
 - straight chain (amylose)
 - branch chain (amylopectin)

Glycogen structure

made of only glucose



- Glucose residue linked α -1,4
- Glucose residue linked α -1,6
- ⊖ Reducing end attached to glycogenin
- ⊙ Nonreducing ends

Digestion of dietary carbohydrates

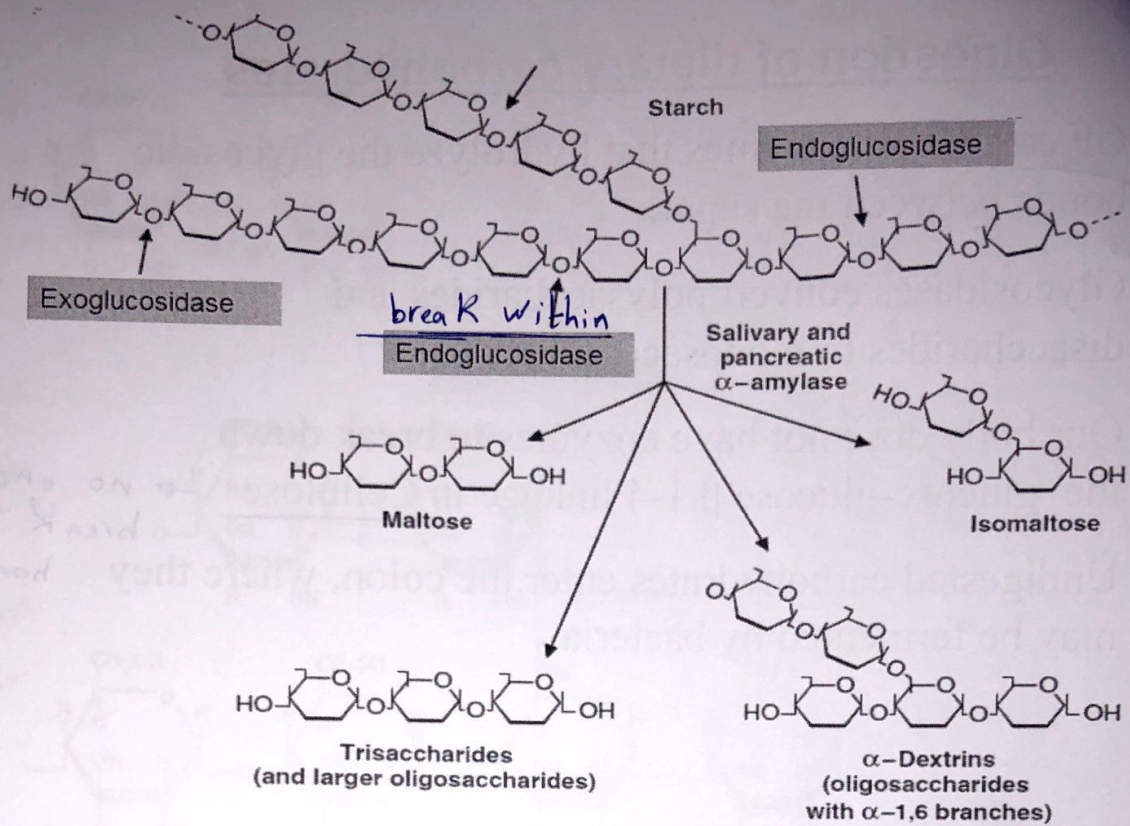
- Glycosidases: enzymes that hydrolyze the glycosidic bonds between the sugars
- Glycosidases convert polysaccharides and disaccharides to monosaccharides
- Our body does not have enzymes to break down the glucose-glucose β 1-4 linkage in Cellulose → no enzyme break this bond
- Undigested carbohydrates enter the colon, where they may be fermented by bacteria

A. Salivary α -amylase

- The digestion of starch begins in the mouth.
- The salivary glands secrete approximately 1 liter of liquid per day into the mouth (pH 6.4 - 7) containing salivary α -amylase.
- α -Amylase is an endoglucosidase, which means that it hydrolyzes internal α -1,4 bonds between glucose residues at random intervals in the polysaccharide chains.
- Requires Cl^- ion for activation with an optimum pH of 6.7 قريبة لـ pH
تأثير اللعاب
- Digestion of starch and glycogen in the mouth gives maltose, isomaltose and α -dextrins
- In the stomach: Carbohydrate digestion stops temporarily, salivary α -amylase may be largely inactivated by the acidity of the stomach.

- If the enzyme starts to break from last molecule

→ Exoglucosidase



• Digestion in the Stomach

- Gastric juice does not contain enzyme for carbohydrate digestion. *due to high acidity*
- Hydrochloric acid in the stomach may hydrolyze sucrose into glucose and fructose.

Digestion in Small Intestine

A. Pancreatic Secretions

- Secretions from the exocrine pancreas (approximately 1.5 L/day) enter the duodenum which contain

- 1- Bicarbonate (HCO_3^-), which neutralizes the acidic pH of stomach contents
 - 2- Pancreatic α -amylase. *similar to salivary amylase*
Pancreatic α -amylase (optimum pH=7.1) also activated by chloride ions, continues to hydrolyze the starches and glycogen, forming the disaccharide maltose, isomaltose, the trisaccharide maltotriose (three glucose molecules linked with α -1,4 glycosidic bonds), and oligosaccharides.
- These oligosaccharides, are usually 4-9 glucose units long linked by α 1-4 and contain one or more α -1,6 branches.
 - α -Amylase has no activity toward sugar containing polymers other than glucose linked by α -1,4 bonds.

→ increase the number of pH to create a proper environment for enzymes

Digestion in Small Intestine

B. Disaccharidases of the intestinal brush-border membrane

1. Glucoamylase

- It has two catalytic sites with similar activities.
- Glucoamylase is an exoglucosidase that is specific for the α -1,4 bonds between glucose residues.
- It begins at the non-reducing end of saccharides
- The glucoamylase is heavily glycosylated with oligosaccharides that protect it from digestive proteases.
- α -amylase split of large polysaccharides molecule and thus supplying new substrate molecules for glucoamylase

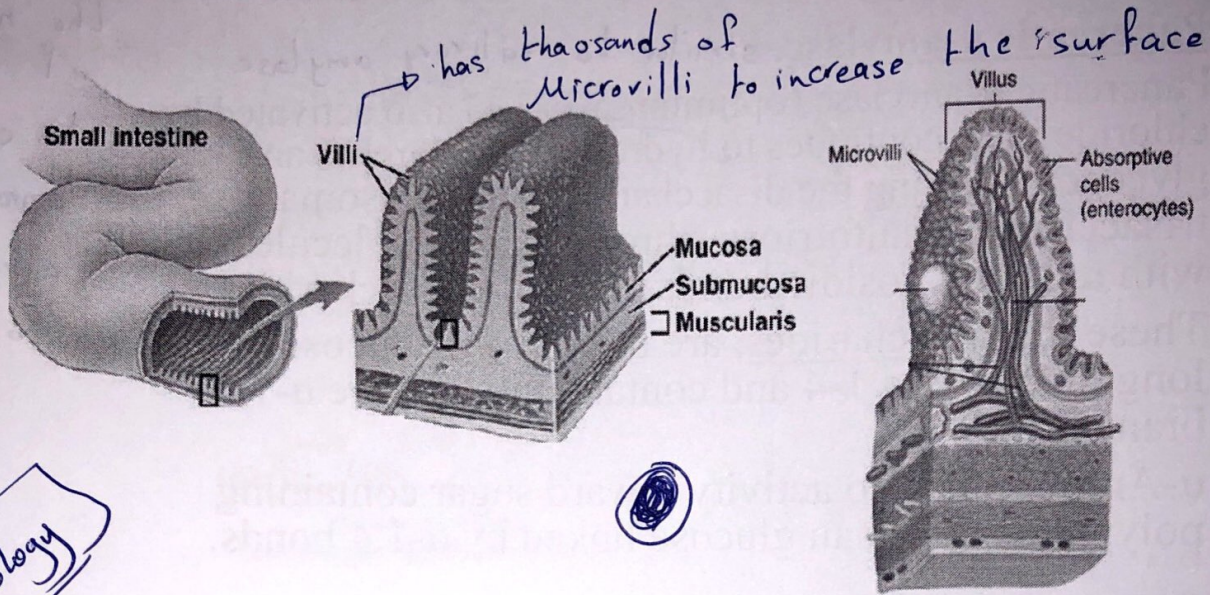
α -amylase of pancreas and saliva functional mainly on polysaccharides (starch) into di or tri saccharides or short chain of oligosaccharides

but glucoamylase breaks the short chain into di or tri saccharides

Location of disaccharide complexes in intestinal villi

2. Suc

- The villi are finger like projections of the mucosa into the lumen of the small intestine.
- A villus has thousands of microvilli.
- Collectively, microvilli known as a "brush border".

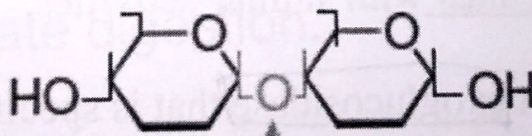


Cardiology
سرونی کا مینل
میراجی

Glucoamylase activity. Glucoamylase is an α -1,4 exoglycosidase, which initiates cleavage at the nonreducing end of the sugar.

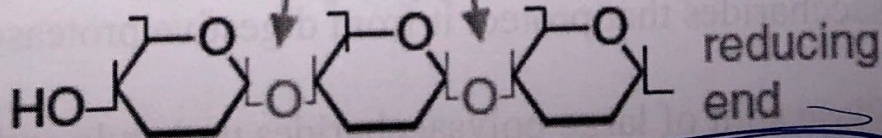
Maltose

α -1,4 bond



maltase activity

1 2



non reducing

reducing end

سرونی کا مینل
Maltotriose

2. Sucrase-isomaltase complex

- Sucrase-isomaltase has two catalytic sites that differs in substrate specificity from the other (Intestinal protease clips it into two separate subunits)

The two catalytic sites are:

(A) The sucrase-maltase site

= Nearly 100% of sucrose hydrolyze (glucose- α -1,2-fructose)

= Maltase activity.

degradation

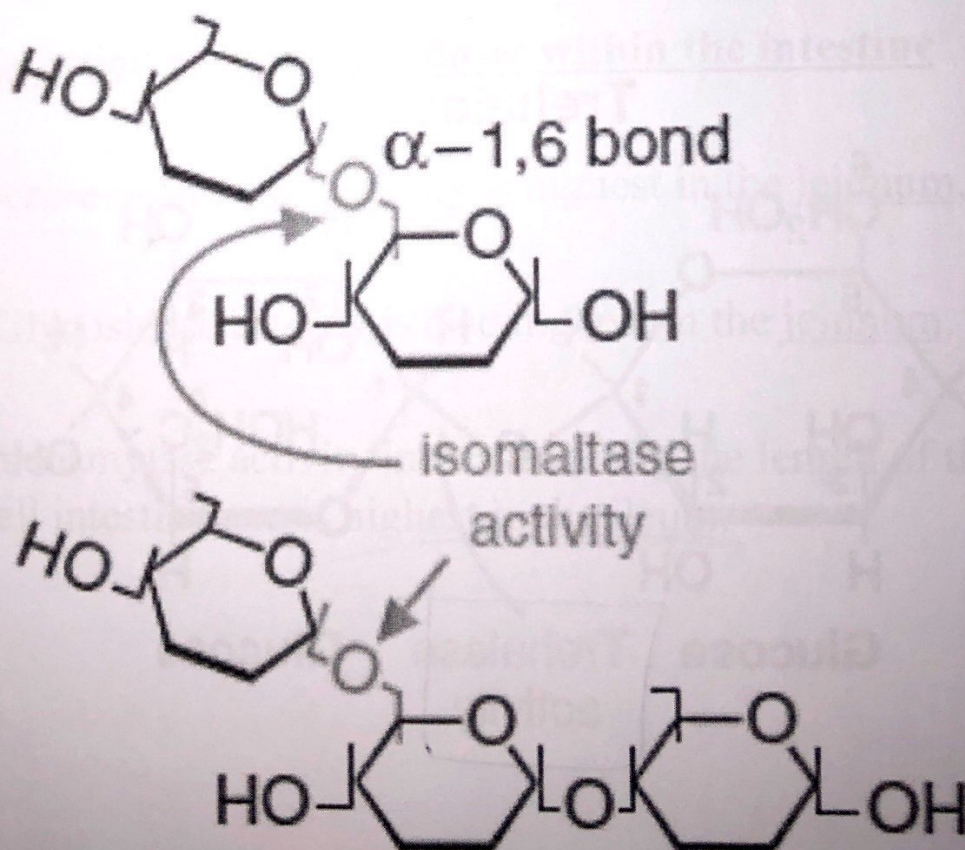
(B) The isomaltase-maltase site

= All α -1,6-glucose-glucose bonds

= Maltase activity.

80% of the maltase is by these complex (maltose: glucose α -1,4 glucose) activity of the small intestine. The remainder of the maltase activity is found in the glucoamylase complex.

20% broken by glucoamylase



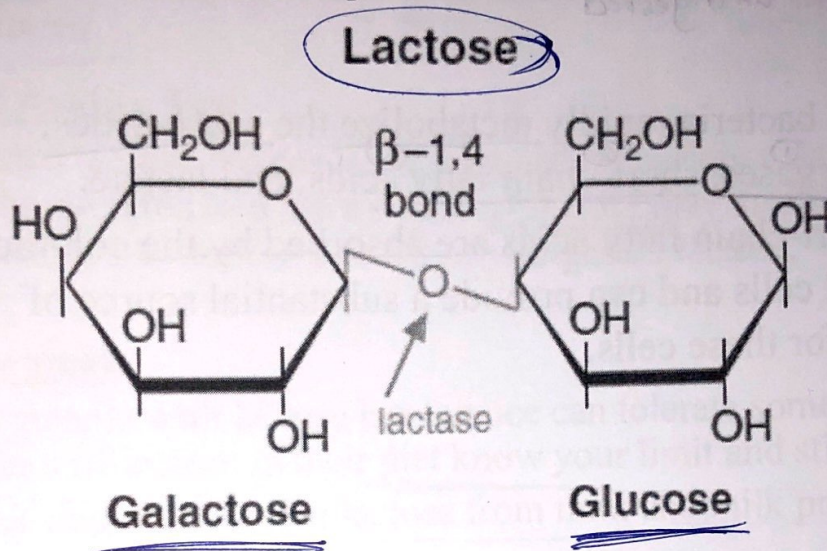
always we takes the function of enzyme by its name.

4. β -glycosidase complex (lactase-glucoylceramidase)

- Has two catalytic sites

1- The lactase catalytic site hydrolyzes the β -bond connecting glucose and galactose in lactose (also splits the β -1,4 bond between some cellulose disaccharides).

2- The β -bond between glucose or galactose and hydrophobic residues ceramide in glycolipids



Location of disaccharidases within the intestine

- ✓ A- Sucrase-isomaltase activity is highest in the jejunum.
- ✓ B- β -Glycosidase activity is also highest in the jejunum.
- ✓ C- Glucoamylase activity increases along the length of the small intestine and is highest in the ileum.

• Metabolism of Sugars by Colonic Bacteria

- Starches high in amylose, or less well hydrated (e.g., starch in dried beans), are resistant to digestion and enter the colon. *undigested*
- Colonic bacteria rapidly metabolize the saccharides, forming ^①gases, ^②short-chain fatty acids, and ^③lactate.
- The short-chain fatty acids are absorbed by the colonic mucosal cells and can provide a substantial source of energy for these cells.

مشان
هيك
تعر
بج
gases
مساكي
beans
(normal case)

Lactose Intolerance not normal (disease)

- The small intestine does not make enough of the enzyme lactase.
- The lactose that is not absorbed is converted by colonic bacteria to lactic acid, methane gas (CH_4), and H_2 gas that result in: abdominal pain, gases, and diarrhea.

Lactase deficiency may be

- (i) **Congenital**: complete deficiency of lactase enzyme since birth (rare).
- (ii) **Acquired**: which occurs later on in life. which include:
 - Primary lactase deficiency** develops over time and begins after about age 2 when the body begins to produce less lactase with a possible genetic link.
 - Secondary** lactase deficiency results from injury to the small intestine, gastrointestinal diseases, including exposure to intestinal parasites.

How is lactose
1. Hydrogen
The person drink

How is lactose intolerance diagnosed?

✓ 1. Hydrogen Breath Test.

The person drinks a lactose-loaded beverage and then the breath is analysed at regular intervals to measure the amount of hydrogen. Normally, very little hydrogen is detectable in the breath, but undigested lactose produces high levels of hydrogen.

✓ 2. Stool Acidity Test

Undigested lactose creates lactic acid and other fatty acids that can be detected in a stool sample. Glucose may also be present in the stool as a result of undigested lactose.

Management:

- 1- Most people with lactose intolerance can tolerate some amount of lactose in their diet know your limit and stick to it.
2. Decreasing or removing lactose from milk and milk products.

- avoid*
- enzyme*
- Glycemic index *time needed of our body to digest the carbs*
 - Not all complex carbohydrates are digested at the same rate within the intestine, and some carbohydrate sources lead to a near-immediate rise in blood glucose levels *and reached the blood*
 - slowly raise blood glucose levels
 - The glycemic index of a food is an indication of how rapidly blood glucose levels rise after consumption.
 - Glucose has the highest glycemic indices (142) with white bread defined as an index of 100.
 - The glycemic response to ingested foods depends not only on the glycemic index of the foods, but also on the fiber and fat content of the food, as well as its method of preparation.

Glycemic Index of Selected Foods, with Values Adjusted to White Bread of 100

Breads	normal	Legumes	
Whole wheat	100	Baked beans (canned)	70
Pumpernickel (whole grain rye)	88	Butter beans	46
Pasta		Garden peas (frozen)	85
Spaghetti, white, boiled	67	Kidney beans (dried)	43
Cereal grains		Kidney beans (canned)	74
Barley (pearled)	36	Peanuts	15
Rice (instant, boiled 1 min)	65	Fruit	
Rice, polished (boiled 10-25 min)	81	Apple	52
Sweet corn	80	Apple juice	45
Breakfast cereals		Orange	59
All bran	74	Raisins	93
Cornflakes	121	Sugars	
Muesli	96	Fructose	27
Cookies		Glucose	142
Oatmeal	78	Lactose	57
Plain water crackers	100	Sucrose	83
Root vegetables		Dairy Products	
Potatoes (instant)	120	Ice cream	69
Potato (new, white, boiled)	80	Whole milk	44
Potato chips	77	Skim milk	46
Yam	74	Yogurt	52

needs more time to digest

because it need less time to digest

• معدة معانات تعرف كم يدوم وقت كحد السكر يصل الدم
 • hypoglycemia = to يتقدر السكر لانه بعد sudden increase في مستوى السكر
 drink

Absorption of sugars

Two mechanisms are responsible for absorption of monosaccharides: active transport and facilitated diffusion.

Features	Facilitated diffusion	Active transport
Concentration gradient	Down the concentration gradient from high to low.	Against a concentration gradient from low to high
Energy expenditure	none	Energy expenditure is in the form of ATP
Carrier protein/ transporter	required	required
Speed	Fast	Fastest mode
Monosaccharides	Fructose and mannose	Glucose and Galactose

Note: Glucose and other monosaccharide's are polar molecule. It cannot pass through lipid bilayer of cell therefore passive diffusion doesn't apply for sugar.

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because it needs less time to digest

• $\frac{d[\text{Glucose}]}{dt} = k[\text{Glucose}]$ sudden increase in glucose drink hypoglycemia

Absorption of sugars

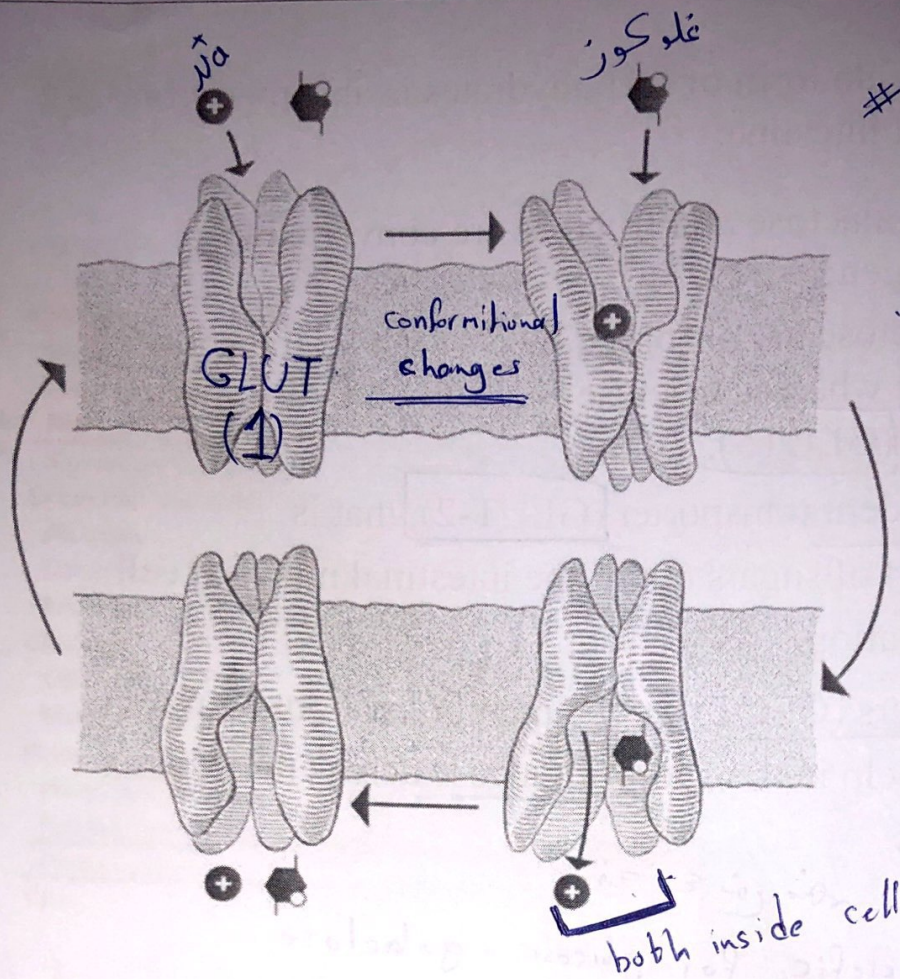
Two mechanisms are responsible for absorption of monosaccharides: active transport and facilitated diffusion.

(Na⁺ ion dependent)

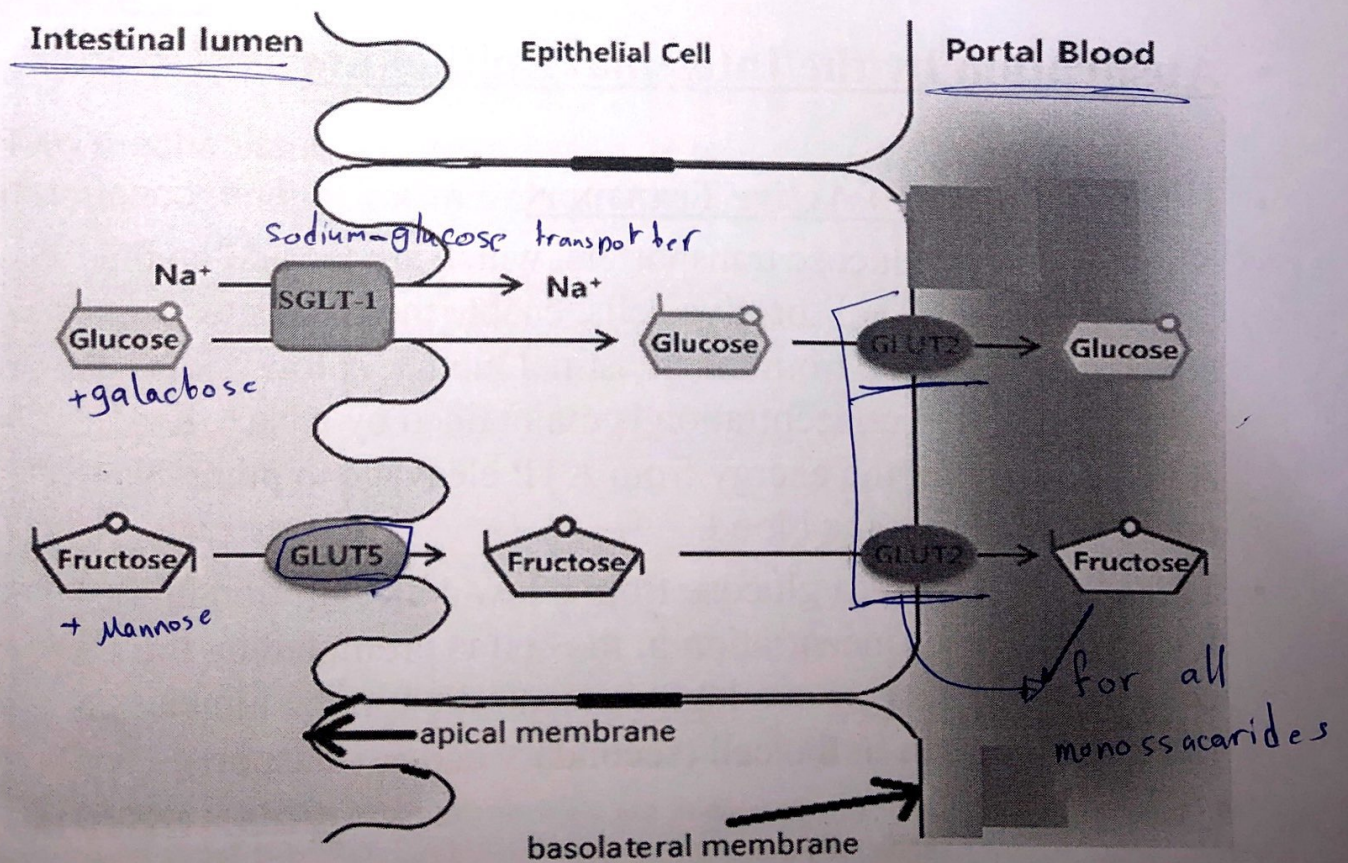
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Speed	Fast	Fastest mode
Monosaccharides	Fructose and mannose	Glucose and Galactose

Note: Glucose and other monosaccharides are polar molecules. It cannot pass through lipid bilayer of cell therefore passive diffusion doesn't apply for sugar. can't pass alone

mono + poly + oligo = monosaccharide



Na^+ binds to the cotransporter, which results in a conformational change that permits glucose to bind to the glucose-binding site. The cotransporter undergoes a new conformational shift, placing Na^+ and glucose near the inner surface of the membrane.



- **2. Facilitative glucose transporters (Na⁺ independent transporter)**
- Glucose moves via the facilitative transporters from the high concentration inside the cell to the lower concentration in the blood without the need of energy. In addition to the Na⁺-dependent glucose transporters, facilitative transporters for glucose also exist on the luminal side of the absorptive cells.
- The various types of facilitative glucose transporters found in the plasma membranes of cells referred to as GLUT 1 to GLUT 14.
- One common structural theme to these proteins is that they all contain 12 membrane-spanning domains.
- Because glucose leaves the intestine via the hepatic portal vein, the liver is the first tissue it passes through. The liver extracts a portion of this glucose from the blood.

they are insulin dependent

Dietary fiber

undigested polysaccharides

- Are components of food cannot be digested by human digestive enzymes they are mainly polysaccharide derivatives and lignan (Noncarbohydrate, polymeric derivatives of phenylpropane).
- There are several kinds of dietary fiber.
- ✓ **Water insoluble fiber:** Cellulose, lignin and hemicellulose are materials that stimulate regular function of the colon.
- ✓ **Water-soluble fiber** Pectins and gums are materials that form viscous gel-like suspensions in the digestive system slowing the rate of absorption of many nutrients, including carbohydrates, and lowering serum cholesterol in many cases.

mainly small intestine

Benefits of fibers

1- Soluble fibers are fermented by bacteria and produce small chain of FA. 10% of our total calories we get from compounds produced by bacterial digestion of substances in our digestive tract.

2- Fiber is thought to "soften" the stool, thereby reducing pressure on the colonic wall and enhancing expulsion of feces this specifically beneficial effect to diverticular disease, in which sacs or pouches may develop in the colon because of a weakening of the muscle and submucosal structures.

△ because the fiber keep water in it

3- Disease prevention for example:

pectins = may lower blood cholesterol levels by binding bile acids = beneficial for diabetes by slowing the rate of absorption of simple sugars and preventing high blood glucose levels after meals. → slow degradation of Pub and their

β-glucan (obtained from oats) reduce cholesterol levels through a reduction in bile acid degradation in the intestine.

absorption is the same

sudden increase in suger ← rate of suger + slow absorption

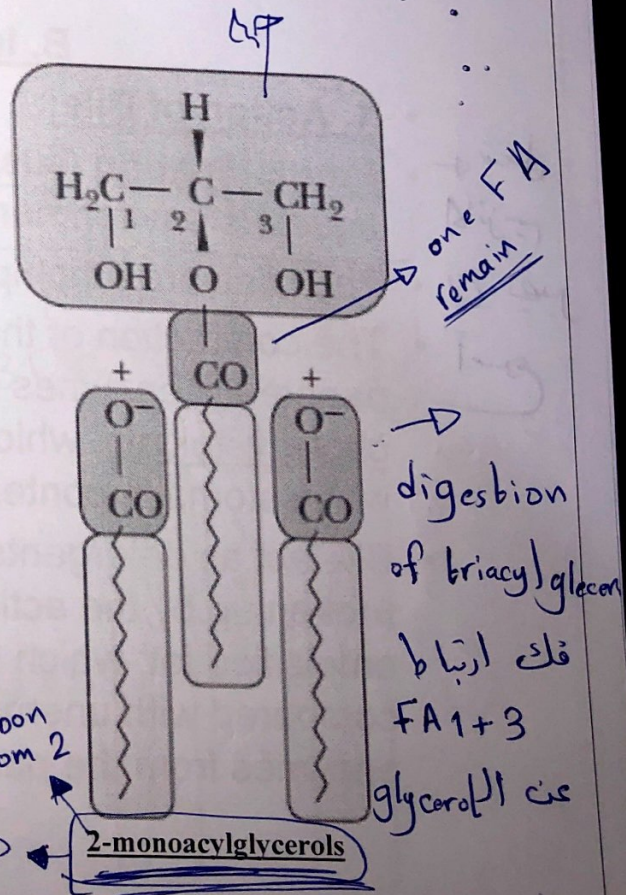
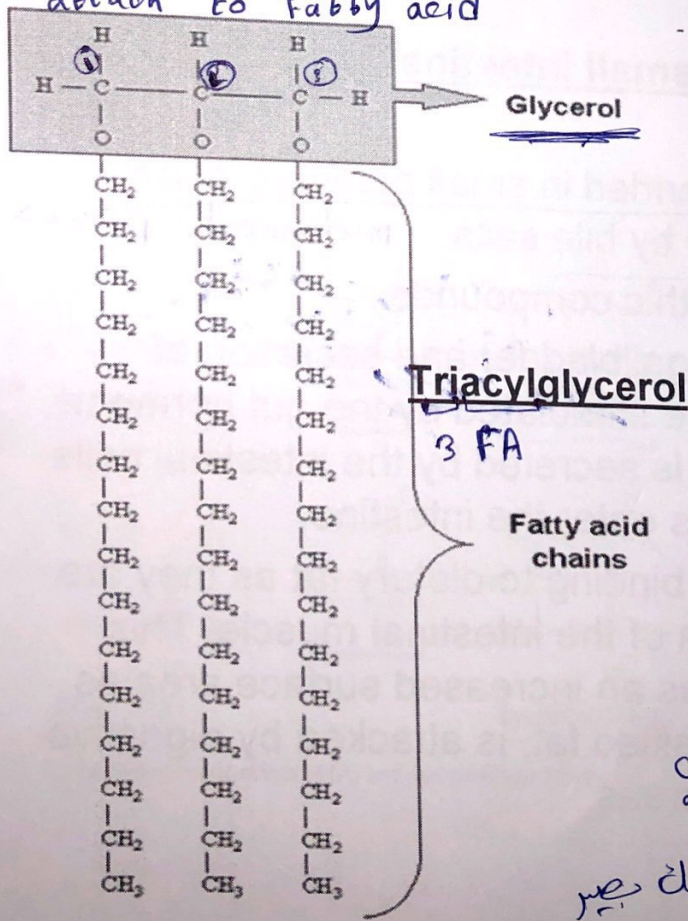
2. Fats

- Functions: (1) Cell structure, (2) fuel storage, (3) hormone all steroid hormones from cholesterol
- Essential fatty acids and nonessential fatty acids
- Nonessential fatty acids can be synthesized in our body essential fatty acids we get them from food We can't synthesize it.
- The essential fatty acids α-linoleic (1) and α-linolenic acid (2) are supplied by dietary plant oils.
- More than 300 different fatty acids are known however only 20-25 of them are widely distributed in nature

Triacylglycerols

- Triacylglycerols are the major fat in the human diet because they are the major storage lipid in the plants and animals that constitute our food supply.
- The remainder of the dietary lipids consists primarily of cholesterol, phospholipids, and free fatty acids.
- Triacylglycerols contain a glycerol backbone to which three fatty acids are esterified.
- The main route for digestion of triacylglycerols involves hydrolysis to fatty acids and 2-monoacylglycerols in the lumen of the intestine. However, the route depends to some extent on the chain length of the fatty acids.

each carbon atom attach to fatty acid



Digestion of fat

A. Mouth and stomach

Limited digestion of lipids occurs in the mouth and stomach because of the low solubility.

1. Lingual lipase produced by cells at the back of the tongue
 2. Gastric lipase produced by stomach
- Same function for both enzymes: they hydrolyze short- and medium-chain fatty acids triacylglycerols (containing 12 or fewer carbon atoms).
 - Lingual and gastric lipases hydrolyse 25-30% of ingested triglycerides (n-3) into diglycerides and free fatty acids.

only hydrolyse one FA in C3

B. In small intestine

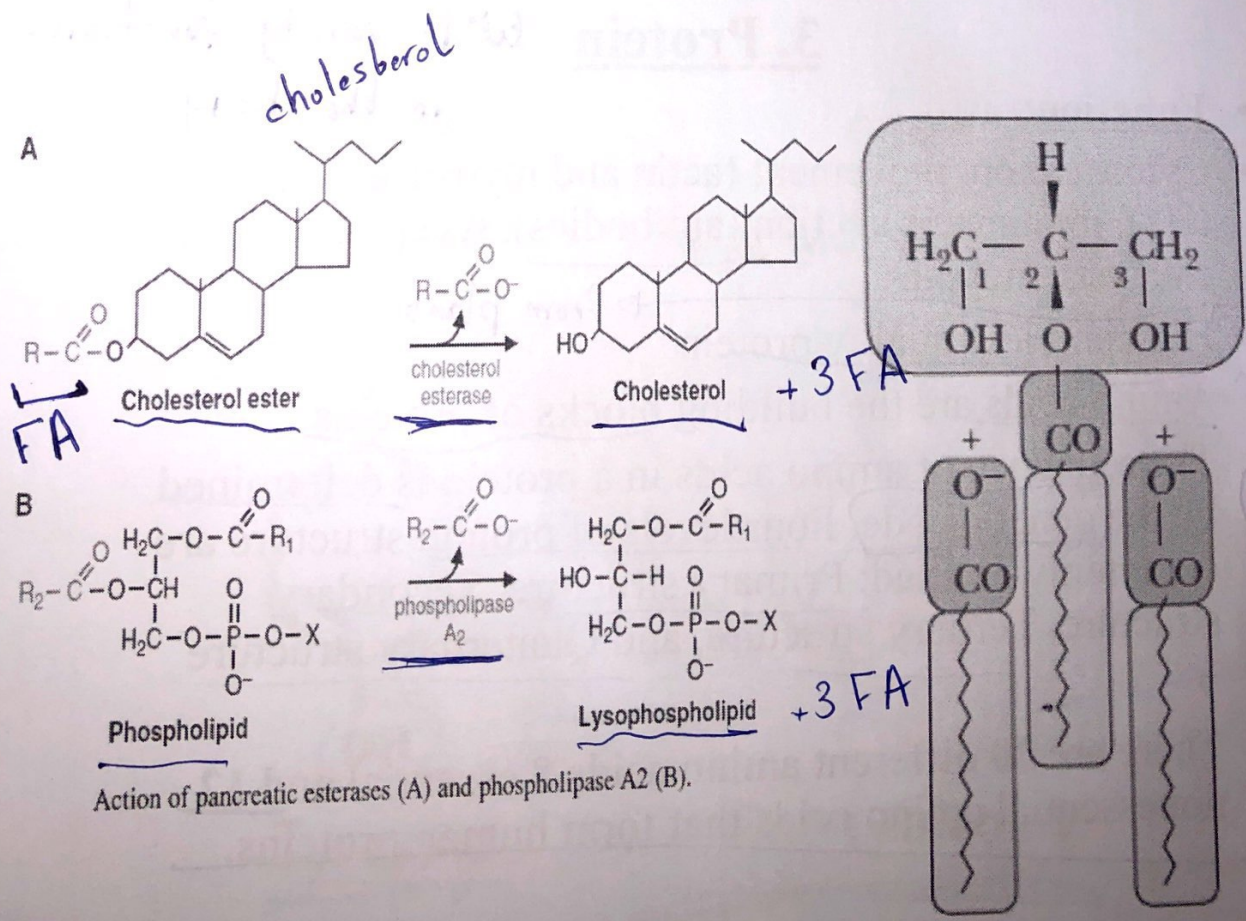
• 1. Action of Bile:

- Emulsification (suspended in small particles in the aqueous environment) by bile salts. *mechanical processes*
- The biles are amphipathic compounds
- The contraction of the gallbladder and secretion of pancreatic enzymes are stimulated by the gut hormone cholecystinin, which is secreted by the intestinal cells when stomach contents enter the intestine.
- Bile act as detergents, binding to dietary fat as they are broken up by the action of the intestinal muscle. This emulsified fat, which has an increased surface area as compared with unemulsified fat, is attacked by digestive enzymes from the pancreas.

شغل
الانزيم
ح بصير
أسرع

2. Action of Pancreatic secretion

- A- Bicarbonate (hormone secretin released from the intestine when acid enters the duodenum) raises the pH of the contents of the intestinal lumen into a range (pH ~ 6.5).
- B- Lipase digests dietary triacylglycerols producing 2 free fatty acids and 2-monoacylglycerol
- C- Colipase it binds to the dietary fat and to the lipase, thereby increasing lipase activity.
- D- Esterases remove fatty acids from compounds such as cholesterol esters (The ester bond is formed between the carboxylate group of a fatty acid and the hydroxyl group of cholesterol.)
- E- phospholipase A2 digests phospholipids to a free fatty acid and a lysophospholipid



• Absorption of digested fats

- Short- and medium-chain fatty acids (C4 to C12) are absorbed directly into intestinal epithelial cells, they enter the portal blood and are transported to the liver bound to serum albumin.

تحتاج ناقل لأنها hydrophobic

- The digested fatty acids and 2-monoacylglycerols are resynthesized into triacylglycerols in intestinal epithelial cells, then packaged in lipoprotein particles chylomicrons because they are insoluble in water, and secreted by way of the lymph into the blood.

synthesized by

ترجع
تكون
سبب
intestinal
epithelium
cells

إذا دخلت بدون ناقل إلى الدم تتعطل blockage
If triacylglycerols directly entered the blood, they would come together, impeding blood flow.

- Chylomicrons transport lipids to adipose tissue, heart, and skeletal muscle. The fatty acids of the chylomicron triacylglycerols are stored mainly as triacylglycerols in adipose cells.

3. Protein

Wide variety functions in the body

- Function:
- Cytoskeleton, movement (actin and myosin), transport (Hb), immune protection (antibodies), receptors and as catalyts enzymes

- High and low quality protein
- Amino acids are the building blocks of proteins.

The sequence of amino acids in a protein is determined by the genetic code. Four levels of protein structure are commonly defined: Primary structure, Secondary structure, Tertiary structure, and Quaternary structure

- There are 20 different amino acids 8 essential and 12 nonessential amino acids that form human proteins.

From animals
(they contain
all essential
Amino acid)

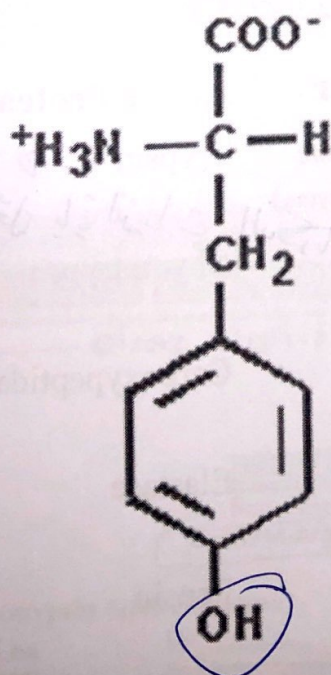
From plants

Conditionally essential amino acids

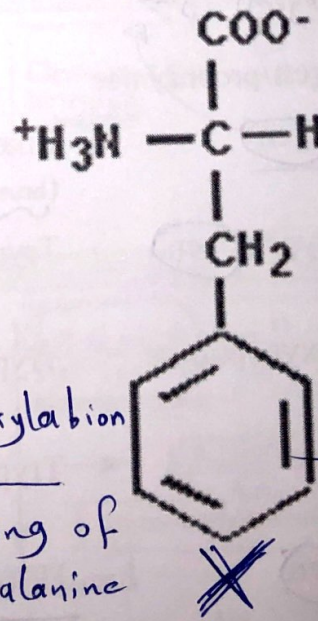
normally they are non essential but become essential under some conditions.

- Children and pregnant women have a high rate of protein synthesis to support growth, and require more of arginine and histidine than their body synthesise.

- Tyrosine is also considered conditionally essential. Tyrosine is synthesized from phenylalanine (by hydroxylation of phenylalanine), and it is required in the diet if phenylalanine intake is inadequate, or if an individual is congenitally deficient in an enzyme required to convert phenylalanine to tyrosine (the congenital disease phenylketonuria).



Tyrosine



Phenylalanine

hydroxylation
of ring of
phenylalanine

لما يكون الفينيل الالانين
في كميات كافية
يكون tyrosine
nonessential
لما اذا كان فيه
نقص يغير essential

proteases ⇒ break down the proteins, very active and powerful (if it work all the time it will digest our body) ⇒ So keep it inactive only if needed

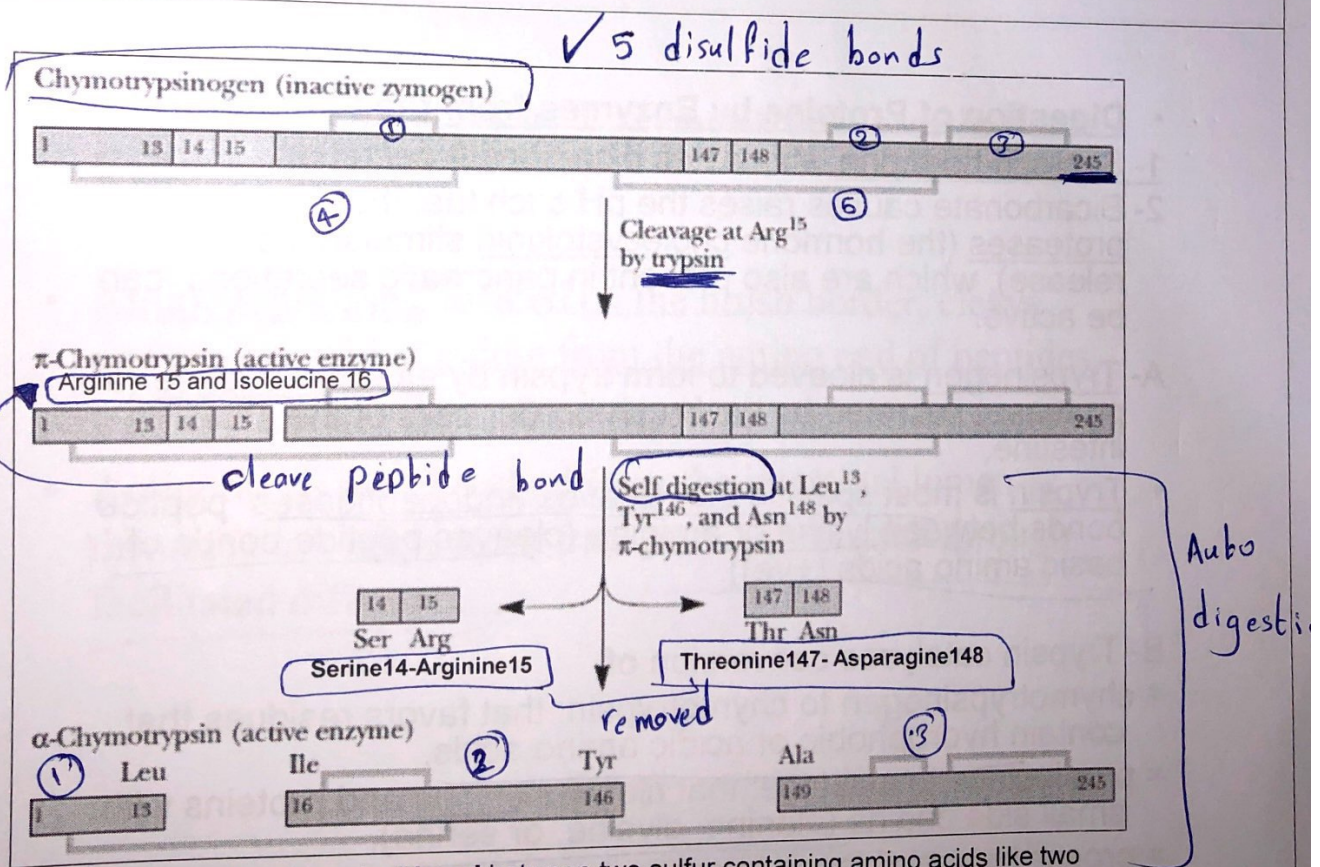
• Zymogens

- Inactive enzymes precursors are called zymogens or proenzymes,
- It only acquire full activity upon specific proteolytic cleavage of one or several of their peptide bonds.
- Zymogen activation by specific proteolysis is an irreversible process.
- Important to switch on processes at the appropriate time and place
- The synthesis of zymogens as inactive precursors prevents them from cleaving proteins prematurely at their sites of synthesis or secretion.

Origin	Zymogen/proenzymes	Activator	Active Protease
<u>Pancreas</u>	Trypsinogen	Enteropeptidase (brush border enzyme)	Trypsin → <i>يسين ينشط</i>
<u>Pancreas</u>	Chymotrypsinogen	Trypsin <i>السكريناس</i>	Chymotrypsin <i>ينذهب ويفعل باقي انزيمات</i>
<u>Pancreas</u>	Procarboxypeptidase	Trypsin	Carboxypeptidase
<u>Pancreas</u>	Proelastase	Trypsin	Elastase
<u>Stomach</u>	Pepsinogen	H ⁺ (pH drop)	Pepsin

زيموجين (لمين)

- **Activation of chymotrypsinogen**
- **Chymotrypsinogen** is a 245-aa cross-linked by five disulfide bonds.
- Chymotrypsinogen is converted to an enzymatically active form called π -chymotrypsin when trypsin cleaves the peptide bond joining Arginine 15 and Isoleucine 16.
- The enzymatically active π -chymotrypsin acts upon other π -chymotrypsin molecules, removing two dipeptides, Serine14-Arginine15 and Threonine147- Asparagine148.
- The end product of this processing pathway is the mature protease α -**chymotrypsin**, in which the three peptide chains.
- A (residues 1 through 13), B (residues 16 through 146), and C (residues 149 through 245), remain together because they are linked by two disulfide bonds, one from A to B, and one from B to C.



A **disulfide bond** is a covalent **bond** between two sulfur containing amino acids like two cysteine molecules

3 polypeptide separated join together by disulfide bond.

• Digestion of Proteins in the Stomach

- ✓ # **Gastrin** hormone (gastric mucosa) stimulates the secretion
- ✓ A- Pepsinogen by the chief cells of the gastric glands.
 - ✓ B- Hydrochloric acid by the parietal cells *high acidity of Hydrochloric acid causes conformational changes in pepsinogen → then Auto digested*
- A- **Pepsinogen** is activated to its active form **pepsin** by acidic gastric juice (pH 1.0 to 2.5) that alters the conformation of so that it can cleave itself, producing the active pepsin.
- **Pepsin** acts as an endopeptidase, cleaving peptide bonds at various points within the protein chain. *unfolding*
- B- Stomach acidity: causes dietary proteins denaturation, this serves to inactivate the proteins and partially unfolds them such that they are better substrates for proteases.
- Smaller peptides and some free amino acids are produced.

• Digestion of Proteins by Enzymes from the Pancreas

- 1- Secretin hormone: stimulates bicarbonate secretion
 - 2- Bicarbonate causes raises the pH such that the pancreatic proteases (the hormone cholecystokinin stimulate its release), which are also present in pancreatic secretions, can be active.
- A- Trypsinogen is cleaved to form trypsin by enteropeptidase (a protease) secreted by the brush-border cells of the small intestine.
- Trypsin is most specific that cleaves endopeptidases peptide bonds between lysine or arginine (cleaves peptide bonds of basic amino acids (+ve)).
- B- Trypsin catalyzes conversion of
- = chymotrypsinogen to chymotrypsin: that favors residues that contain hydrophobic or acidic amino acids.
 - = proelastase to elastase: that cleaves elastin and proteins with small side chains (alanine, glycine, or serine).
 - = procarboxypeptidases to carboxypeptidases.