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CARBOHYDRATE CHEMISTRY Mucopolysaccharides by Dr. Shuzan Ali Mohammed Assist. Prof. of

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MUCOPOLYSACCHARIDES (Glycosaminoglycans GAGs)

I. Neutral mucopolysaccharides:

- Formed of protein & polysaccharides.
- Contain <u>acetyl hexosamines but no uronic acid.</u>
- Present in mucous secretion.

II. Acidic mucopolysaccharides: A. Non-sulfated

B. Sulfated



Heterogeneous (mucopolysaccharides GAGs)						
I. Neutral	II. Acidic					
NANA in tissue of vertebrate & bacteria	A. Non- sulfated	Hyaluronic a.				
Bl. gp subs. <u>(L-fucose</u> is	B. sulfated	Heparin				
important constituent) Gonadotrophins &		Heparan sulphate				
thyrotrophic H		Chondoritin sulfate				
a1 & a2 globulins		Keratan sulphate				
Ovalbumin		-				
Fibrinogen		Dermatan sulphate				



N.B. L-fucose is deoxyhexose at C6 (C6H12O5) NANA: Pyruvate + mannosamine

Mucopolysaccharides (negatively-charged)

- Most GAG are present extracellular except heparin
- They act as lubricant and cushion for other tissues as they absorb large amount of water
- On compression of GAG; water is squeezed out and they occupy smaller volume. When the compression is released they return to their original volume. This property is called <u>resilience</u> of synovial fluid and vitrous humor of the eye.

Acidic MPS

Formed of (repeated disaccharide units): 1- amino-sugar acids OR amino- sugars 2- Uronic acid (glucuronic or iduronic) OR monosaccharide Linked by glycosidic bond



II. Acidic mucopolysaccharides A. Non-sulfated mucopolysaccharides: <u>1.Hyaluronic acid</u>

•Repeating units of <u>N-acetyl glucosamine & B-glucuronic a.</u> Functions:

- Forms the <u>cement substance</u> between tissues.
- Present in synovial fluid (lubricant facilitates joint movement)
- Makes cartilage <u>compressible</u>.
- Makes **ECM loose** (by the ability to attract H_2O).
- Permits <u>cell migration</u> during wound repair &

morphogenesis.

GlcNAc
$$\xrightarrow{\beta_{1,4}}$$
 GlcUA $\xrightarrow{\beta_{1,3}}$ GlcNAc $\xrightarrow{\beta_{1,4}}$ GlcUA $\xrightarrow{\beta_{1,3}}$



(**N.B.**)

- It facilitates cell migration; being produced in increased amount by tumor cells; so facilitates migration through ECM & spread of tumor.
- <u>Hyaluronidase</u> secreted by <u>certain bacteria</u> causes destruction of this cement subs. (hyaluronic a) so help spread of infection <u>(spreading factor)</u>.
- <u>Hyaluronidase</u> is present in <u>acrosomal cap of</u> <u>sperm</u> & invades the tissues of the ova causing destruction of hyaluronic a. & its fertilization.
- Morphogenesis: cell differentiation into tissues & organs in the embryo



A. Sulfated mucopolysaccharides:
<u>1-Heparin</u>: repeating units of:
<u>Sulfated glucosamine &</u> <u>Sulfated glucuronic a. (or L-iduronic a.)</u>
linked by <u>α-1,4 glycosidic bond</u>.
Formed by <u>mast cells (intracelluar), located along the</u> blood vessel wall in many tissues like heart, lung, liver, kidney, skin & spleen.

Its concentration in blood is very low.

N.B. Iduronic acid is C5 epimer of glucouronic acid (differ in the position of –OH at C5)



Functions of heparin 1. Anticoagulant: It activates antithrombin III & It Inhibits blood clotting factors II, VII, IX & X. 2. Plasma clearance from lipids: It activates lipoprotein lipase that digests plasma lipids. (heparin & lipoprotein lipase are clearing factors)



2- Heparan sulphate:

It differs from heparin in the amount of uronic a. & the sulphate attached to glucosamine (more glucuronic a. but less sulphated glucosamine).
It is a component of <u>ECM</u> in the form of proteoglycans. It has a role in:

Cell-cell interaction &

2. Cell membrane receptors.

 $\begin{array}{c} \text{6-Sulfate} \\ \text{IdUA} \xrightarrow{\alpha_{1,4}} & \text{GlcN} \xrightarrow{\alpha_{1,4}} & \text{GlcUA} \\ & & & \\ & & & \\ & & & \\ & & & \\ 2\text{-Sulfate} & & SO_3^- & \text{or Ac} \end{array}$

3- Chondroitin sulfate: repeating units of: <u>N-acetyl galactosamine & B-glucuronic a. linked</u> by **<u>B-1,3 bond.</u>** Types: 3 (A, B & C) **1- Chondroitin sulfate A:** sulfate ester gp of N- acetyl galactosamine at C4 \rightarrow Chondroitin-4-sulfate A at $C6 \rightarrow Chondroitin-6$ -sulfate C Chondroitin itself is a minor component of ECM but its sulfate ester (A & C) are major component of <u>cartilage</u>, bone, cornea & other connective tissues. 2- Chondroitin sulfate B: It yields upon hydrolysis L-iduronic a. instead of D-glucuronic a.



4- Keratan sulphate: repeating units of:
Galactose and N-acetyl glucosamine linked together by β- bond.
No uronic acid.
Present in cornea (to make it transparent),

cartilage and tendons.



5- Dermatan sulphate: repeating units of:
L-iduronic acid and N-acetyl galactosamine linked together by α-1,3 bond.
Present in blood vessels, heart, cornea, sclera & skin. It maintains the shape of sclera

> IdUA ^{α1,3}→ GalNAc | | 2-Sulfate 4-Sulfate



Points	Hyaluronic a	Heparin	Heparan S	Chondroitin S	Keratan S	Dermatan S
Amino S.	Glucosamine	Glucosamine	Glucosamine	Galactosamine	Glucosamine	Galactosamine
Uronic a	Glucuronic a	Glucuronic + Iduronic	Glucuronic + Iduronic	Glucuronic + Iduronic	Galactose (No uronic a)	Iduronic a
Sulfate	Absent	Present	Present	Present	Present	Present
Bonds	β-1,3 & β- 1,4	α-1,4	α-1,4	β-1,3	β-1,3 & β- 1,4	α-1,3
Sites	 S.C. tissues Ovum Wall Synovial fluid 	-CT <u>Mast cells</u> (liver, spleen, kidney, bone marrow) but least conc. in blood.	EXTRA- CELLULAR Matrix (ECM)	 Matrix of cartilage Tendons Ligaments Bone 	- <u>Cornea</u> . -Cartilage -Tendons	-Blood vessels -Heart. - Cornea, - <u>Sclera</u> - Skin.
Function	Protective for tissues	Anticoagulant Lipid clearance from plasma	-Cell-cell interaction -Cell membrane receptors	Supportive	Supportive Transparency of cornea	Supportive Maintains the shape of sclera



N.B. <u>When proteins are connected to acidic</u> <u>mucopolysaccharides $\rightarrow \rightarrow$ proteoglycans</u>

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