Lecture 23 (11/10/25)

TODAY

•Reading: Ch7; 251-254, 258-

260, 241-250

•Homework: #21, #22

NEXT

Ch8; 294-296, 263-269 Reading:

Ch24; 885-890

Homework: #23

Carbohydrates

- A. Definition
- - 2. Derivatives: the chemistry of
 - carbohydrates
 - a. Oxidation

 - b. Reduction
 - i. C1/C2 ii. Other carbons
 - c. Ester formation d. Amino sugars
 - 3. Polymerization

 - a. The Glycosidic Bond
 b. Non-covalent bonds in macro-molecular
 - c. Disaccharides
- D. Oligosaccharides
 - 1. Glycoproteins & glycolipids
 - 2. O-linked
 - 3. N-linked
 - 4. Sequence determination-ABO
- E. Polysaccharides
 - 1. Polymers of glucose
 - 2. Polymers of disaccharides

11-11-2025



BU Staff Holiday, not Faculty or Students.

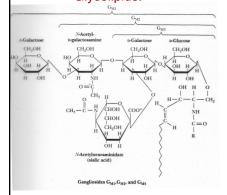
Oligosaccharides

Carbohydrates

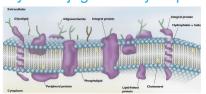
Oligosaccharides:

Here is an oligosaccharide that we will encounter when we study LIPIDS; sphingoglycolipids, or for short:

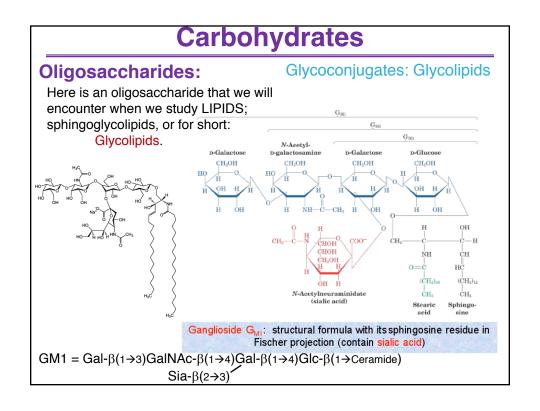
Glycolipids.

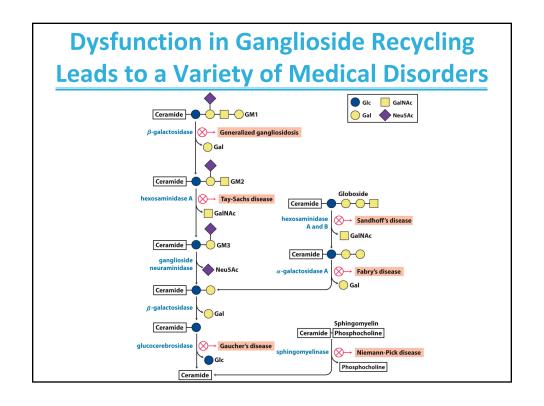


Glycoconjugates: Glycolipids



 $GM1 = Gal-\beta(1\rightarrow 3)GalNAc-\beta(1\rightarrow 4)Gal-\beta(1\rightarrow 4)Glc-\beta(1\rightarrow Ceramide)$ $Sia-\beta(2\rightarrow 3)$





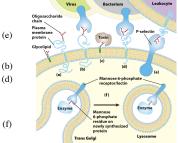
Oligosaccharides:

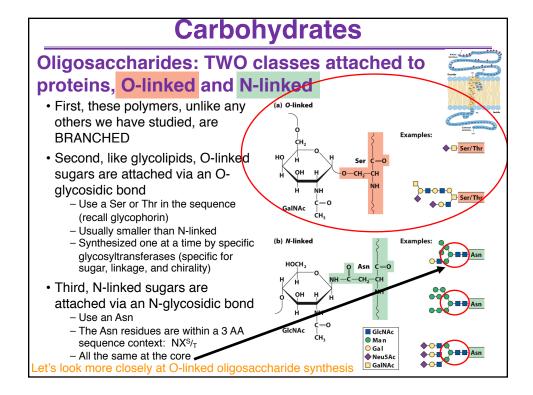
Glycoconjugates: Glycoprotein

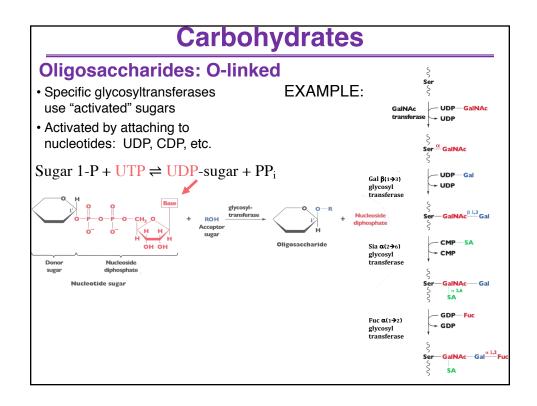
When you attach an oligosaccharide to a protein: Glycoproteins.

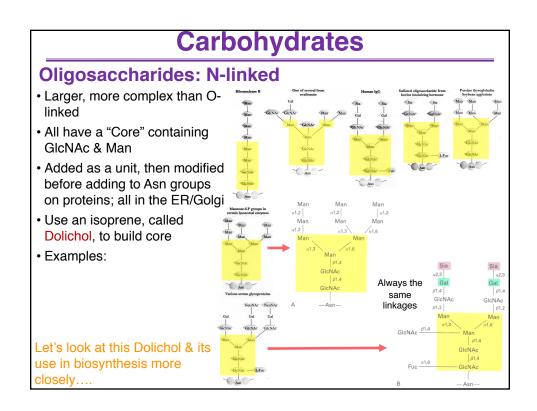
- A protein with small oligosaccharides attached:
 - Carbohydrate is attached via its anomeric carbon to amino acids on the protein.
 - · Common connections occur at Ser, Thr, and Asn.
 - About half of mammalian proteins are glycoproteins.
 - Generally, bacteria do not glycosylate their proteins.
 - Carbohydrates play role in protein-protein recognition. (a)
 - Viral proteins are heavily glycosylated; this helps evade the immune system.
- Proteins whose role is to bind specific carbohydrates/oligosaccharides = Lectins (or selectins)
 - Lectins important for many biological functions
 - Recruitment of leukocytes to sites of inflammation
 - · Sperm-egg recognition
 - · Virus-target cell interaction
 - Attachment of flora (microbiome) in gut
 - · Nervous system development
 - · Serum-protein turnover (sialic acid)
 - · Targeting proteins to lysosomes for degradation (Man) (f

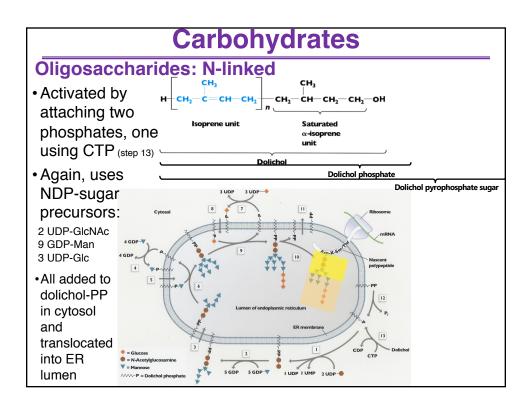
How are these sugars attached?

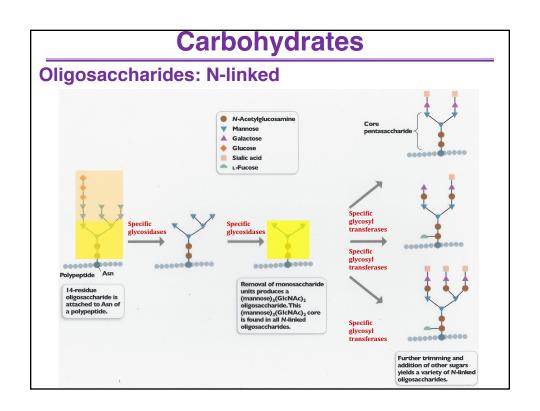










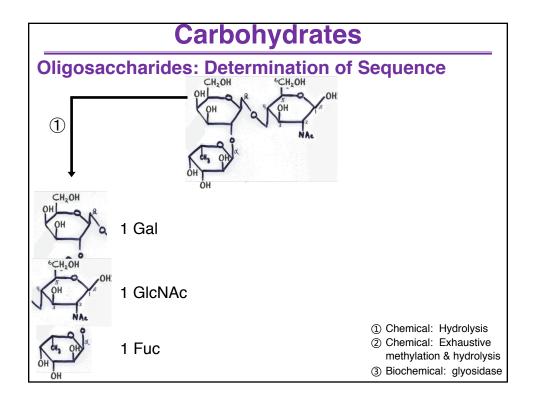


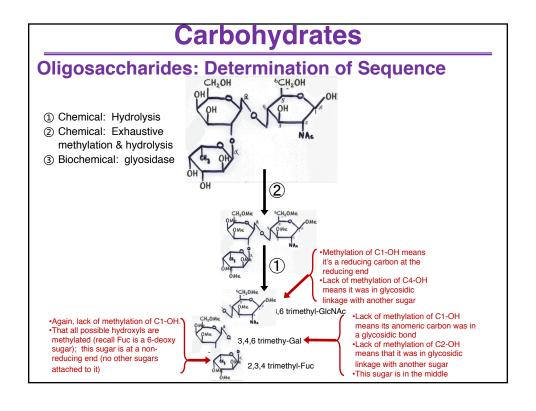
Oligosaccharides: Determination of Sequence

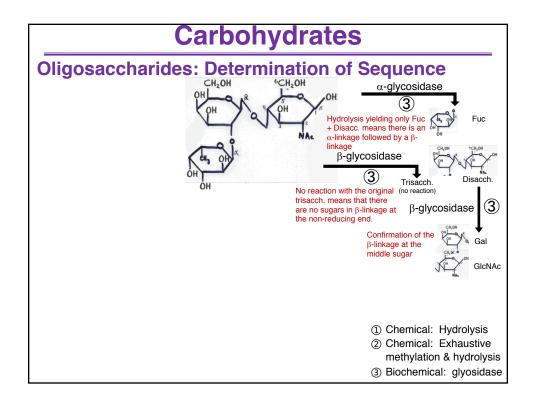
- Whole different problem compared to proteins and nucleic acids.... Its branched!!
- Moreover, a given residue can have several (and stereo-specific) ways of attaching to a neighboring residue.
- Need to use a combination of methods:
 - Chemical
 - Hydrolysis & chromatography to identify sugars
 - Exhaustive methylation & hydrolysis, then chromatography to identify what positions were not methylated
 - Biochemical
 - Use of enzymes that stereo-specifically hydrolyze glyosidic bonds (from the non-reducing end)

EXAMPLE: First, just like protein sequencing, you need to purify glyco-protein or lipid. Lets say we isolate the glycolipid from a person's RBC's who is O-positive. Treat it with a ceramidase to hydrolyze the lipid from the sugar.

Second, take an aliquot and just hydrolyze (like what was done for amino acid analysis). This gets the composition and stoichiometry.

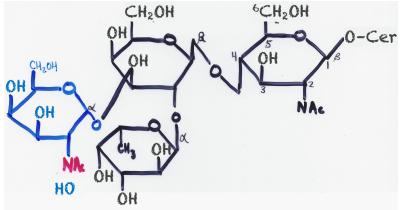






Oligosaccharides: Determination of Sequence

In vertebrates, ganglioside carbohydrate composition determines blood groups.



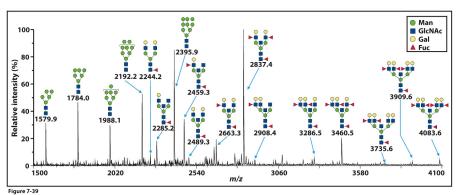
L-Fuc $\alpha(1\rightarrow 2)$ -D-Gal $\beta(1\rightarrow 4)$ D-GlcNAc β -Ceramide O blood group

D-Gal $\alpha(1\rightarrow 3)$ D-GalNAc $\alpha(1\rightarrow 3)$

B blood group

A blood group

Glycoconjugates: Analysis



Lehninger Principles of Biochemistry, Seventh Edition © 2017 W. H. Freeman and Company

Polysaccharides

Carbohydrates

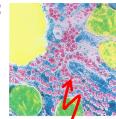
Polysaccharides

- The majority of natural carbohydrates are usually found as large polymers.
- These polysaccharides can be:
 - homopolysaccharides (one monomer unit)
 - heteropolysaccharides (multiple monomer units)
 - linear (one type of glycosidic bond)
 - branched (multiple types of glycosidic bonds)
- Polysaccharides do not have a defined molecular weight.
 - This is in contrast to proteins because, unlike proteins, no template is used to make polysaccharides.
 - Polysaccharides are often in a state of flux; monomer units are added and removed as needed by the organism.

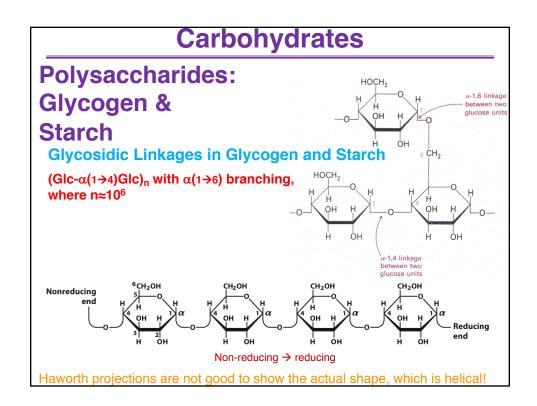
Polysaccharides: Polymers of Glucose

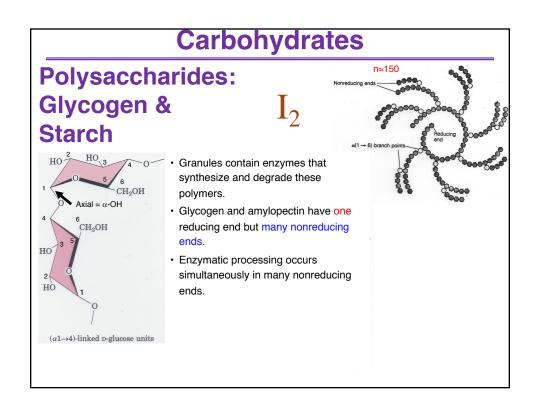
Homopolymers of Glucose:

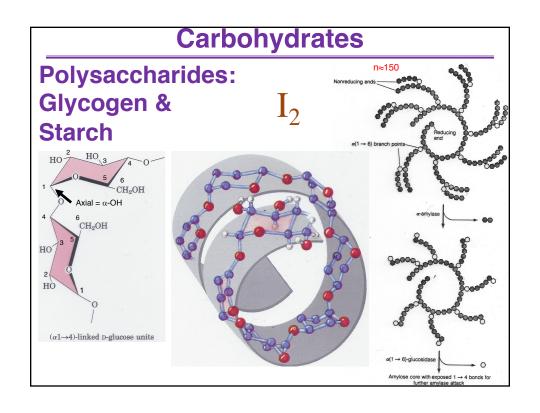
- Starch
- Glycogen
- Cellulose
- Chitin

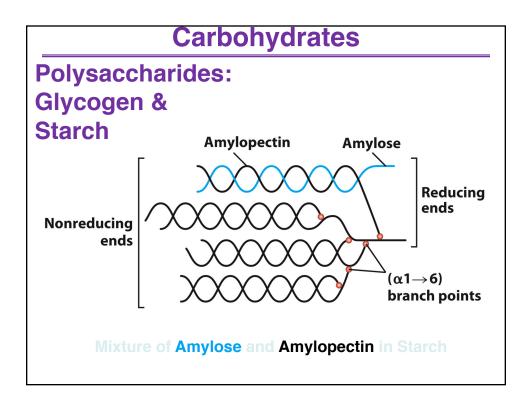


- Glycogen and Starch are the main storage polysaccharides for energy.
 - Glycogen and starch are insoluble due to their high molecular weight and often form granules in cells.
 - Molecular weight reaches several millions (~200 x 10⁶)(can see in microscope).
- Glycogen is a branched homopolysaccharide of glucose.
 - Glucose monomers form $\alpha(1 \rightarrow 4)$ linked chains.
 - There are branch points with $\alpha(1 \rightarrow 6)$ linkers every 8–12 residues.
- Starch is a mixture of two homopolysaccharides of glucose.
 - Amylopectin is like glycogen, but the branch points ($\alpha(1 \rightarrow 6)$ linkages) occur every 24–30 residues.
 - Amylose is an unbranched polymer of $\alpha(1 \rightarrow 4)$ linked residues.









Polysaccharides:

Homopolymers of Glucose:

- Starch
- Glycogen
- Cellulose
- Cellulose is a linear homopolysaccharide of glucose.
 - Glucose monomers form β(1 → 4) linked chains (no branching).
 - Hydrogen bonds form between adjacent monomers.
 - There are additional H-bonds between chains.
 - Structure is now tough and water insoluble.
 - It makes up almost 50% of plant mass; in cell walls.
 - Cotton is nearly pure fibrous cellulose.



Polysaccharides:

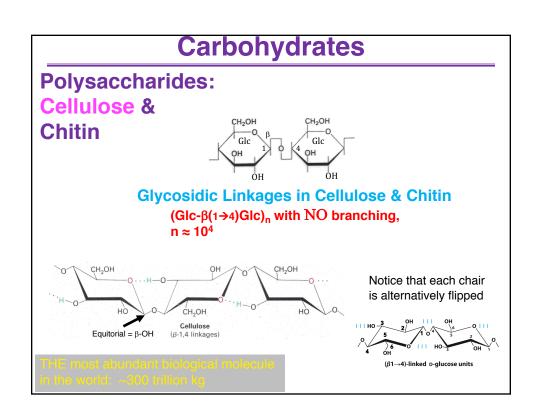
Homopolymers of Glucose:

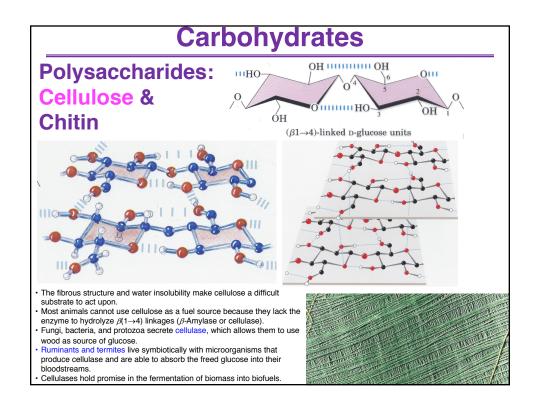
- Starch
- Glycogen
- Cellulose
- Chitin

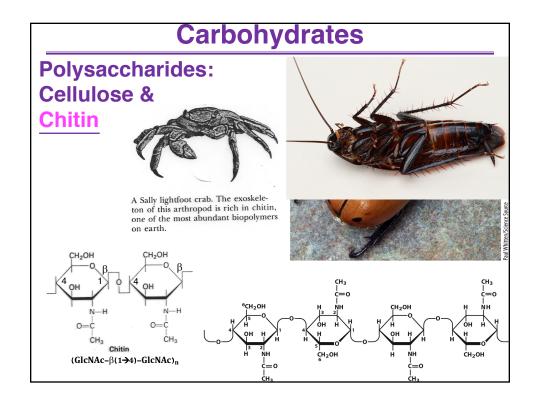


 Chitin is a linear homopolysaccharide of Nacetylglucosamine (GlcNAc).

- N-acetylglucosamine monomers form $\beta(1 \rightarrow 4)$ -linked chains.
- forms extended fibers that are similar to those of cellulose
- hard, insoluble, cannot be digested by vertebrates
- structure is tough but flexible, and water insoluble
- found in cell walls in mushrooms and in exoskeletons of insects, spiders, crabs, and other arthropods



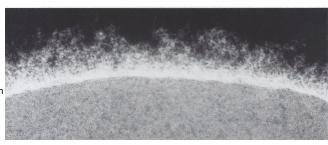


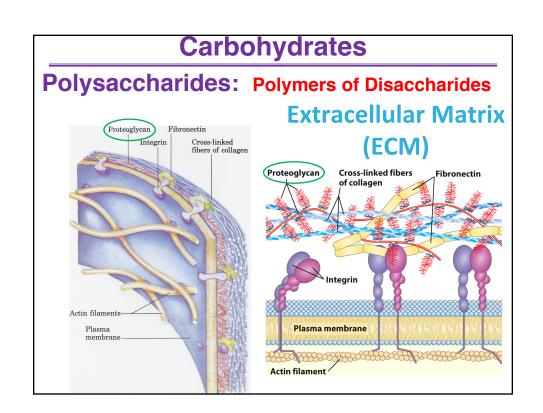


Polysaccharides: Polymers of Disaccharides

First, need to describe the Extracellular Matrix (ECM)

- · Material outside the cell in most animal cells
- · Strength, elasticity, and physical barrier in tissues (varies tremendously)
- · Main components
- -proteoglycans
- -collagen & elastin fibers
- Proteoglycans
- Different glycosaminoglycans are O-linked to the "core protein."
- Linkage from anomeric carbon of xylose to serine hydroxyl
- Our tissues have many different core proteins; aggrecan is the best studied.





Polysaccharides: Polymers of Disaccharides Glycosaminoglycans

(the carbohydrate part of proteoglycans)

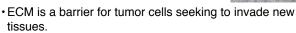
- Linear polymers of repeating disaccharide units (sugarX-sugarY)_n
- One monomer (sugarX) is either sugar acid or Gal
 - uronic acids (C6 oxidation)
 - Most have sulfate esters
- One monomer (sugarY) is either:
 - N-acetyl-glucosamine (GlcNAc) or N-acetyl-galactosamine (GalNAc)
 - Also sulfate esters
- Extended hydrated molecule
 - Negatively charged
 - minimizes charge repulsion
- Forms meshwork with fibrous proteins to form extracellular matrix
 - connective tissue
 - lubrication of joints
- Form huge $(M_r > 2 \cdot 10^8)$ noncovalent aggregates (Hyaluronan and Aggrecan).
 - They hold a lot of water (1000× its weight) and provide lubrication.
 - Very low friction material
 - Covers joint surfaces: articular cartilage
 - o reduced friction & load balancing

Carbohydrates

Polysaccharides: Polymers of Disaccharides

Extracellular Matrix (ECM)

- Main components
- -proteoglycans
- -collagen & elastin fibers
- Proteoglycans
- -Different glycosaminoglycans are O-linked to the "core protein."



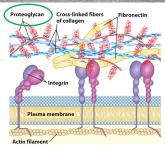
Some tumor cells secrete heparinase that degrades ECM.

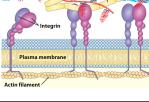
Cosmetics



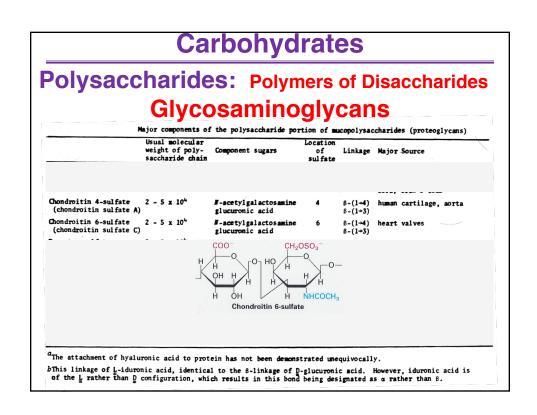




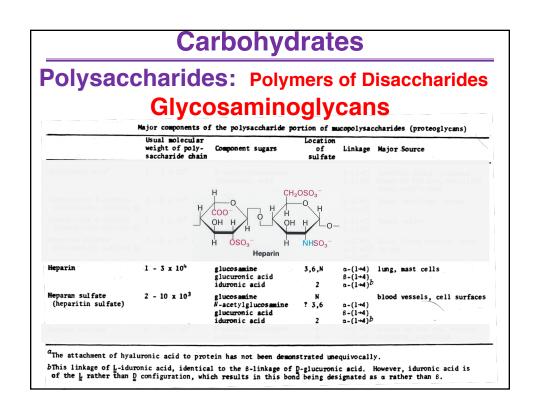


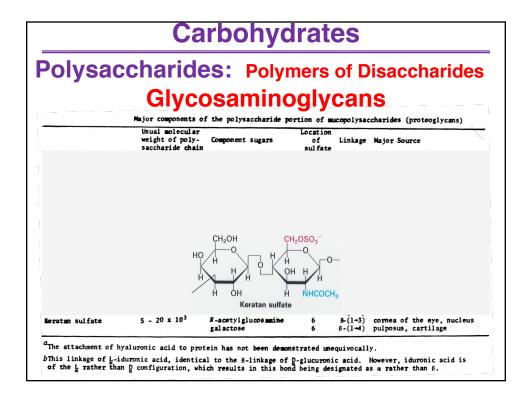


Carbohydrates Polysaccharides: Polymers of Disaccharides Glycosaminoglycans Major components of the polysaccharide portion of mucopolysaccharides (proteoglycans) Usual molecular weight of poly- component sugars sucharide chain sufface Hyaluronic acid 1 - 3 x 106 #-acetylglucosamine glucuronic acid 5-(1-4) synovial fluid, vitreous 8-(1-3) humor of the eye, umbilical cord, cock's comb Chool Hyaluronate The attachment of hyaluronic acid to protein has not been demonstrated unequivocally. This linkage of L-iduronic acid, identical to the 8-linkage of D-glucuronic acid. However, iduronic acid is of the L rather than 2 configuration, which results in this bond being designated as a rather than 8.

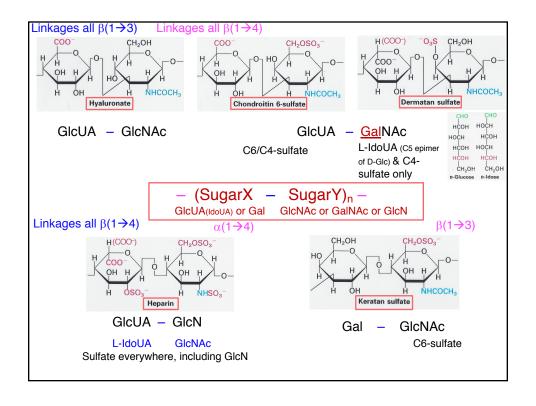


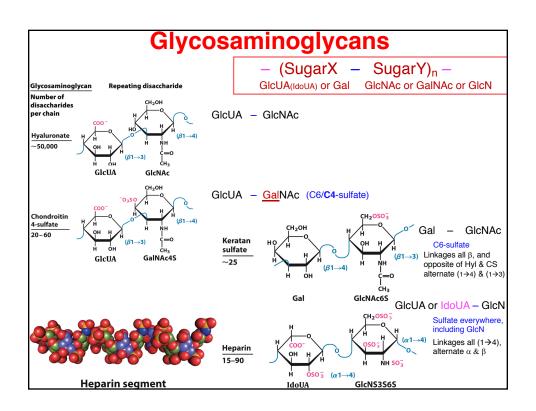
Carbohydrates Polysaccharides: Polymers of Disaccharides Glycosaminoglycans Major components of the polysaccharide portion of mucopolysaccharides (proteoglycans) Usual molecular weight of polysaccharide portion of mucopolysaccharides (proteoglycans) Location of binkage Major Source Dermatan sulfate (chondroitin sulfate B) A sectylgalactosamine of sulfate sulfate sulfate sulfate Component sugars of binkage Major Source A sectylgalactosamine of sulfate sulfat

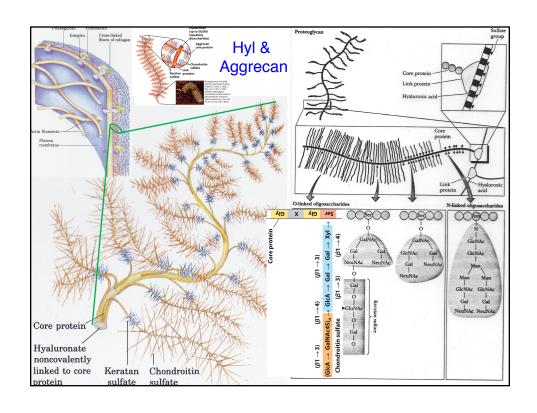


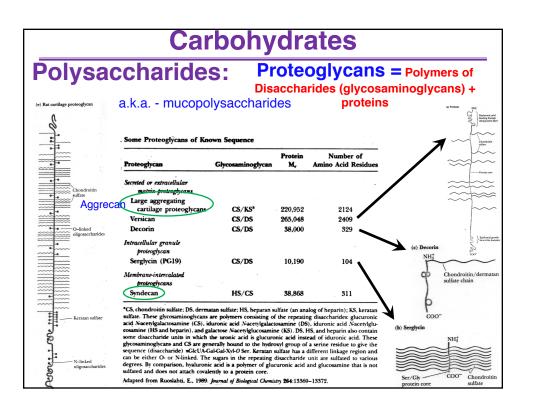


Carbohydrates Polysaccharides: Polymers of Disaccharides							
Glycosaminoglycans							
nal molecular ight of poly- charide chain	Component sugars	Location of sulfate		Major Source			
- 3 x 10 ⁶	N-acetylglucosamine glucuronic acid	•		synovial fluid, vitreous humor of the eye, umbilical cord, cock's comb			
5 x 104	W-acetylgalactosamine glucuronic acid	4	β-(1+4) β-(1+3)	human cartilage, aorta			
- 5 x 10 ⁴	N-acetylgalactosamine glucuronic acid	6	β-(1+4) β-(1+3)	heart valves			
5 x 10 ⁴	N-acetylgalactosamine iduronic acid glucuronic acid	4	β-(1-4) α-(1-3) ^b β-(1-3)	skin, blood vessels, heart valves			
3 x 10 ⁴	glucuronic acid iduronic acid	3,6,N 2	α-(1+4) β-(1+4) α-(1+4) ^b	lung, mast cells			
10 x 10 ³	glucosamine N-acetylglucosamine glucuronic acid	N ? 3,6	α-(1+4) β-(1+4)	blood vessels, cell surfaces			
20 x 10 ³	iduronic acid N-acetylglucosamine galactose	2 6 6	8-(1+3)	cornea of the eye, nucleus pulposus, cartilage			
	components of the land solecular ght of poly- charide chain 3 x 10 ⁶ 5 x 10 ⁴ 5 x 10 ⁴ 5 x 10 ⁴ 10 x 10 ³	Glycosamino components of the polysaccharide por nal molecular ght of poly-charide chain 3 x 106 5 x 104 5 x 105 8 -acetylgalactosamine glucuronic acid glucuronic acid glucuronic acid glucuronic acid iduronic acid iduronic acid iduronic acid glucuronic acid glucuronic acid iduronic acid glucuronic acid iduronic acid glucuronic acid iduronic acid glucuronic acid iduronic acid increase in acid	components of the polysaccharide portion of musual molecular ght of poly-charide chain 3 x 106 5 x 104 5 x 105 6 y-acety galactosamine of glucuronic acid glucuronic acid glucuronic acid glucuronic acid iduronic acid iduronic acid iduronic acid iduronic acid glucuronic acid glucuronic acid iduronic acid iduronic acid iduronic acid iduronic acid iduronic acid glucuronic acid iduronic acid iduronic acid glucuronic acid iduronic acid glucuronic acid iduronic acid glucuronic acid iduronic acid iduronic acid 2 20 x 103 8-acety glucosamine of 6	Components of the polysaccharide portion of mucopolysaccharide portion of mucopolysaccharide thain Tomponents of the polysaccharide portion of mucopolysaccharide chain Component sugars Of Linkage sulfate S x 10 th S-acetylglucosamine glucuronic acid glucuronic aci			









Polysaccharides:

Proteoglycans = Polymers of Disaccharides (glycosaminoglycans) +

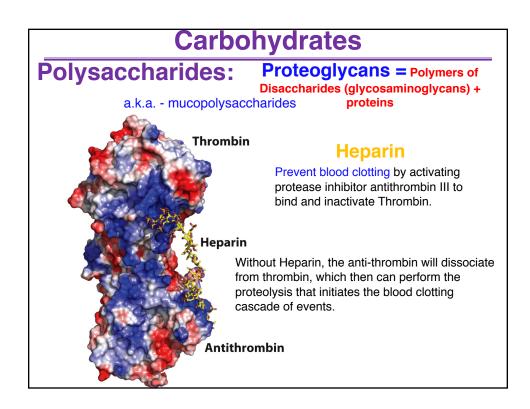
a.k.a. - mucopolysaccharides

proteins

- Some integral membrane proteins are proteoglycans.
 - svndecans
- Other integral membrane proteins are receptors for extracellular proteoglycans.
 - Integrins (which then interact with Actin)
- These proteins link cellular cytoskeleton to the ECM and transmit signals into the cell to regulate:
 - cell growth cell mobility

 - apoptosis
 - wound healing
- In chondrocytes, huge amounts of cartilage proteoglycans are produced (Aggrecan)

 - Cushioning
 - Due to reversible hydration
- Heparin is linear polymer, 3-40 kDa.
 - Heparan sulfate is heparin-like polysaccharide but attached to proteins
 - Highest density of negative-charge of any biomolecule
 - Prevent blood clotting by activating protease-inhibitor antithrombin III
 - Binding to various cells regulates development and formation of blood vessels.
 - Modulates growth and development by binding growth-factor receptors
 - Important role in cancer metastasis
 - Can also bind to viruses and bacteria and decrease their virulence



Primer	Type ^a	Repeating unit ^b	Size (number of monosaccharide units)	Roles/significance
Starch				Energy storage: in plants
Amylose Amylopectin	Homo- Homo-	$(\alpha 1 \rightarrow S4)$ Glc, linear $(\alpha 1 \rightarrow S4)$ Glc, with $(\alpha 1 \rightarrow S6)$ Glc branches every 24–30 residues	50-5,000 Up to 10 ⁶	
Glycogen	Homo-	(α1→S4) Glc, with (α1→S6) Glc branches every 8–12 residues	Up to 50,000	Energy storage: in bacteria and animal cells
Cellulose	Homo-	(β1 → S4) Glc	Up to 15,000	Structural: in plants, gives rigidity and strength to cell walls
Chitin	Homo-	(β1 → S4) GlcNAc	Very large	Structural: in insects, spiders, crustaceans, gives rigidity and strength to exoskeletons
Peptidoglycan	Hetero-; peptides attached	4)Mur2Ac(β1→S4) GleNAc(β1	Very large	Structural: in bacteria, gives rigidity and strength to cell envelope
Hyaluronan (a llycosaminoglycan)	Hetero-; acidic	4)GlcA (β1 → S3) GlcNAc(β1	Up to 100,000	Structural: in vertebrates, extracellular matri of skin and connective tissue; viscosity and lubrication in joints