By Dr. Gouse Mohiddin Shaik

- Introduction
- The gastrointestinal (GI) tract is a complex system performing multiple biological functions which are anatomically distributed
- Site for food processing and absorption
- Largest immune organ also
- At the interface with external environment constitutes barrier against ingested foreing materials like microbes, toxins
- Like with many tissues, succumbs to diseases
- Several biomarkers available

The GI tract

The GI tract

(gastrointestinal tract)

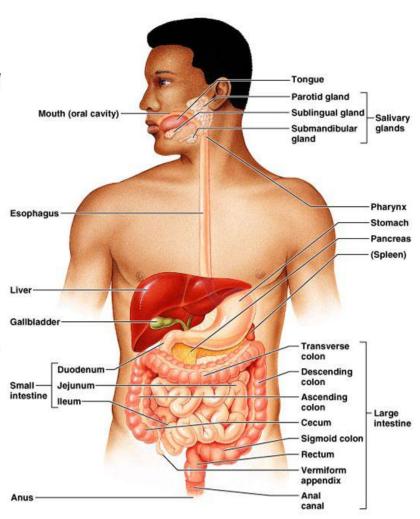
The muscular alimentary canal

- Mouth
- Pharynx
- Esophagus
- Stomach
- Small intestine
- Large intestine
- Anus

The accessory digestive organs

Supply secretions contributing to the breakdown of food

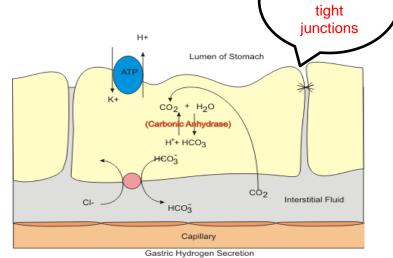
- Teeth & tongue
- Salivary glands
- Gallbladder
- Liver
- Pancreas



- Introduction
- Sampling methods of GI tract diseases
 - Feces analysis many diseases
 - Biopsy cancers
 - Imaging ulcers

- Digestive Biomarkers
- Elastase 1
 - This is a pancreatic enzyme
 - Secreted as zymogen (inactive enzyme)
 - Activated by trypsin in duodenum
 - Remains undegraded during its transit through intestine
 - So serves as very good marker for the function of pancrease
 - Especially useful marker in acute pancreatitis, and pancreatic insuffiency

- Digestive Biomarkers
- Hypochlorohydria VEGF in saliva
 - Characterized by low to no acid production in stomach
 - Leads to improper digestion and absorption
 - VEGF plays role in preventing H+ leaking back thus maintaining acidic environment
 - Less VEGF in saliva indicate disease

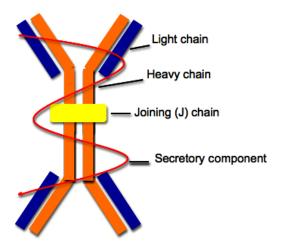


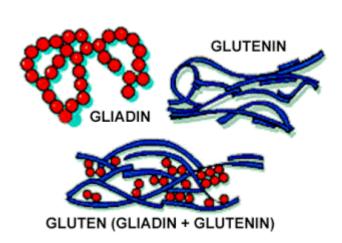
VEGF

maintain

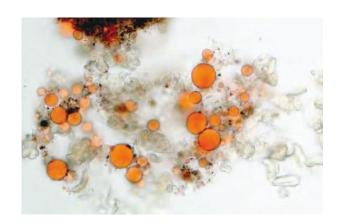
- Digestive Biomarkers
- Triglycerides
 - High levels of triglycerides in feces indicate problems with fat digestion
 - Could result from pancreatic insuffiency, insufficient bile

- Immune Biomarkers
- Fecal IgA
 - IgA found mainly in body secretions
 - Increased slgA is found in conditions like inflammation
 - Specific sIgA like anti-gliadin IgA indicate gluten intolerance



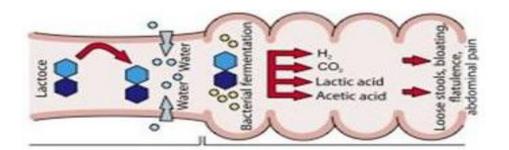


- Absorption biomarkers
- Long chain fatty acids, total fat, cholesterol
 - Healthy GI tract absorbs, malabsorption will result in elevation in stools
- Vegetable fibers and muscle fibers
 - Poor digestion and absorption results in appearance of muscle fibers in feces





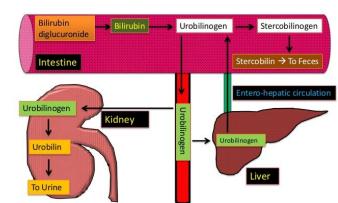
- Absorption biomarkers
- Lactose intolerance
 - Disaccharides are converted in to monosaccharides
 - Lactase required for digestion of lactose
 - Lactase is limited in humans
 - Results in lactose intolerance
 - Leads to production of gas, pain, loose stools....



- Absorption biomarkers
- Steatorrhoea
 - Presence of fat in stools
 - Because of defective fat absorption
 - Defective fat absorption also leads to defective absorption of fat dissolved vitamins
 - Vitamin D and Vitamin K deficiency especially



- Other biomarkers
- Color
 - Abnormal color may be because of excess intake of pigmented food
 - Brown color because of action of intestinal bacteria to produce stercobilinogen
 - Red color because of blood could indicate many conditions like
 - Cancer
 - Ulcer
 - Inflammation
 - Injury



- Other biomarkers
- Fecal occult (hidden) blood test (FBOT)
- Frequently used test is gualac based FBOT

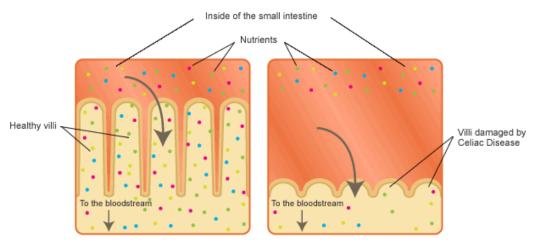
$$\begin{array}{c} \text{Pseudoperoxidase} \\ \text{Hemoglobin} + \text{H}_2\text{O}_2 + \text{guaiac} & \longrightarrow \text{oxidized guaiac} + \text{H}_2\text{O} \\ \text{(colorless)} & \text{(Blue color)} \end{array}$$

 Annual test recommended for early detection of colon cancer



- Other biomarkers
- Neutrophils
- Neutrophils are found in feces during infection with bacteria like salmonella, shigella, yersinia and E. Coli
- Toxin mediated or viral mediated diarrhea do not cause appearance of neutrophils
- Can be detected by imaging and staining pro0cedures

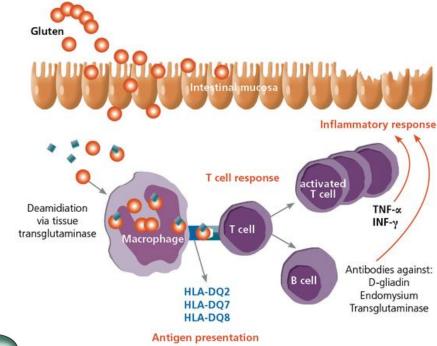
- Other biomarkers
- HLA-DQ varients in celiac disease (predisposing)
- Mainly a autoimmune disorder
- Sensitivity to gluten
- Effects the villi of small intestine, resulting in less abosrption
- Chronic condition may turn in to cancer



 A. In a healthy person, nutrients get absorbed by villi in the small intestine and go into the bloodsteam.

B. In a person with Celiac Disease, the villi have been damaged by inflammation, so fewer nutrients pass into the bloodstream.

- Other biomarkers
- HLA-DQ varients



Chromosome 6

Long arm
Short arm

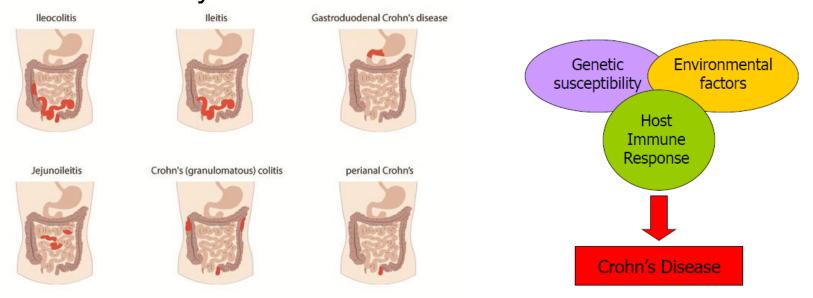
HLA region

Class II

DP DQ DR B C A

HLA Complex

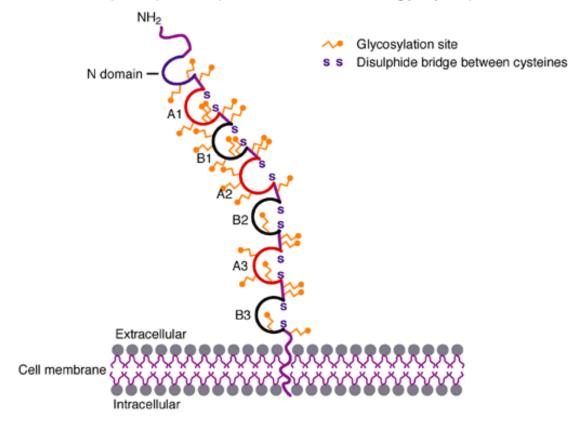
- Other biomarkers
- Crohn's disease
- A type of inflammatory bowel disease
- Characterized by abdominal pain, fever, diarrhoea....
- Still not clear why this disease occurs
- Collective effect of genetics, environment and lifestyle choices may result in disease



- Other biomarkers cancer markers
- Mostly non-specific
- Unpredictable and changes from case to case
- CEA Carcino enbryogenic antigen
- A cell adhesion glycoprotein
- Well studied marker for colorectal cancer
- Produced by fetus, no production after birth
- Some cancers tend to activate the gene and expression level increases
- Can be tracked in blood

Other biomarkers – cancer markers

b Structure of CEA protein (the 70-kD protein becomes 180 kD when glycosylated)



Schematic representation of the human carcinoembryonic antigen (CEA) gene and protein

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Next class

- Next class.....
 - Biomarkers of renal diseases