#### <section-header>

### Absorptive Capability of the Small Intestine

- Length: Approximately 6 meters.
- Surface Area: Enhanced by villi and microvilli, providing a vast area for nutrient absorption. <u>Lybrate</u>

### A Maldigestion vs. Malabsorption

| Aspect     | Maldigestion                          | Malabsorption  |
|------------|---------------------------------------|--|
| Definition | Impaired breakdown of nutrients.      | Defective mucosal uptake and transport of adequately digested nutrients, including |
|            |                                       | vitamins and trace elements.   |
| Example    | Lactase deficiency leading to lactose | Celiac disease affecting the absorption of   |
|            | intolerance.                          | multiple nutrients.  |

# Malabsorption Syndrome

- Definition: A clinical term encompassing defects during the digestion and absorption of food nutrients by the gastrointestinal tract.
- Types:
- Isolated Malabsorption: Impairment of a single nutrient component (e.g., lactose intolerance).
- Global Malabsorption: Diffuse disorders affecting absorption of almost all nutrients (e.g., Celiac disease, Crohn's disease).

#### Pathophysiology: Phases of Digestion & Absorption

- 1. Luminal Phase (Digestion):
- Pancreatic Insufficiency: Chronic pancreatitis, cystic fibrosis, post-surgical states leading to decreased lipase and protease activity.
- Gastric Hypersecretion (Zollinger-Ellison Syndrome): Inactivation of pancreatic enzymes due to low pH.
- Post-Gastrectomy: Inadequate mixing of nutrients, bile, and pancreatic enzymes.
- 2. Mucosal Phase (Absorption):
- Mucosal Damage (Villous Atrophy):
- Celiac disease.
- Intestinal lymphoma.
- Crohn's disease.
- Eosinophilic enteritis.
- Amyloidosis.
- Small intestinal bacterial overgrowth (SIBO).
- Giardiasis.
- Whipple's disease.
- Tropical sprue.
- Viral gastroenteritis.
- Intestinal tuberculosis.
- NSAIDs, Olmesartan.3. Post-Absorptive Phase
- 3. Post-Absorptive Phase (Lymphatic Transport):
- Lymphatic Obstruction:
- Congenital: Intestinal lymphangiectasia.
- Acquired: Whipple disease, lymphoma, tuberculosis, congestive heart failure, constrictive pericarditis, radiation therapy, retroperitoneal fibrosis.

### **W** Clinical Manifestations of Malabsorption

| Symptom/Sign                  | Mechanism   |
|-------------------------------|---|
| Weight loss/Malnutrition      | Anorexia, malabsorption of nutrients.                               |
| Diarrhea                      | Impaired absorption or secretion of water and electrolytes; colonic |
|                               | fluid secretion secondary to unabsorbed bile acids and fatty acids. |
| Flatus                        | Bacterial fermentation of unabsorbed carbohydrates.                 |
| Glossitis, Cheilosis          | Deficiency of iron, vitamin B12, folate, and vitamin A.             |
| Abdominal Pain                | Bowel distention or inflammation, pancreatitis.                     |
| Bone Pain                     | Calcium, vitamin D malabsorption, protein deficiency,               |
|                               | osteoporosis.   |
| Tetany, Paresthesia           | Calcium and magnesium malabsorption.                                |
| Weakness                      | Anemia, electrolyte depletion (particularly potassium).             |
| Azotemia, Hypotension         | Fluid and electrolyte depletion.                                    |
| Amenorrhea, Decreased Libido  | Protein depletion, decreased calories, secondary hypopituitarism.   |
| Anemia                        | Impaired absorption of iron, folate, vitamin B12.                   |
| Bleeding                      | Vitamin K malabsorption, hypoprothrombinemia.                       |
| Night Blindness/Xerophthalmia | Vitamin A malabsorption.  |
| Peripheral Neuropathy         | Vitamin B12 and thiamine deficiency.                                |

#### 🚽 Diarrhea & Steatorrhea

- Diarrhea:
- Most common symptomatic complaint.
- Defined as an increase in stool mass, frequency, or fluidity, typically greater than 200 g per day.
- Steatorrhea:
- Result of fat malabsorption.
- Hallmark: Passage of pale, bulky, malodorous stools that float and are difficult to flush.
- Presence of floating oil droplets in the toilet.

#### Weight Loss & Fatigue

- Weight Loss:
- Common and may be pronounced.
- Patients may compensate by increasing caloric consumption, masking weight loss.
- More likely in diffuse diseases like celiac disease and Whipple disease.
- Fatigue
- Often due to anemia and nutrient deficiencies. Medscape

#### Flatulence & Abdominal Distention

- Caused by bacterial fermentation of unabsorbed food substances, releasing gases like hydrogen and methane.
- Leads to uncomfortable abdominal distention and cramps.

### Edema

- Hypoalbuminemia from chronic protein malabsorption or loss into the intestinal lumen causes peripheral edema.
- Extensive lymphatic obstruction (e.g., intestinal lymphangiectasia) can cause protein loss.
- Severe protein depletion may lead to ascites.

# Anemia

- Types:
- Microcytic: Iron deficiency, often seen in celiac disease.
- Macrocytic: Vitamin B12 deficiency, common in ileal involvement or resection (e.g., Crohn's disease).

#### Metabolic Bone Disease

- Vitamin D deficiency can cause osteopenia or osteomalacia.
- Bone pain and pathologic fractures may be observed.
- Calcium malabsorption can lead to secondary hyperparathyroidism.

# Carbohydrate Malabsorption

- Etiology:
- Lactase deficiency (most common).
- Post-intestinal resection.
- Mucosal diseases.
- Post-infectious gastroenteritis.
- Dietary changes (e.g., Eastern to Western diet).
- Workup:
- Stool Osmolality: Normally equals plasma osmolality (~290 mOsm/kg).
- Stool Osmotic Gap: 100 mOsm/kg indicates osmotic diarrhea.
- Stool pH: <6 suggests fermentation.</li>
- Lactase DNA Assay: Genetic testing for lactase deficiency.
- Hydrogen Breath Test: Increased hydrogen indicates carbohydrate malabsorption.

# Celiac Disease (Continued)

# Epidemiology

- Primarily affects whites of northern European ancestry.
- Prevalence in at-risk groups:
- 1 in 22 → first-degree relatives
- 1 in 39 → second-degree relatives
- 1 in 56 → symptomatic patients
- 1 in 300–500 → general population

# Pathogenesis

- Genetic factors:
- Strong intrafamilial occurrence
- HLA-DQ2 and/or DQ8 genes (in ~36%)
- Associated with chromosome 15q26 (linked to T1DM)
- Non-HLA loci → inherited as autosomal recessive

### Classifications

| Туре      | Description  |
|-----------|--|
| Classic   | Villous atrophy + malabsorption symptoms + improvement on        |
|           | gluten-free diet   |
| Latent    | Recovered as child with gluten-free diet, remained silent, then  |
|           | developed CD again later   |
| Potential | Positive serology/genetics but normal biopsy; risk of developing |
|           | CD later   |

# ${\cal S}$ Association with Type 1 Diabetes

- Approximately 5% of patients with type 1 diabetes develop celiac disease.
- Common initial findings include:
- Unpredictable blood glucose levels.
- Recurrent hypoglycemia.
- Poor glycemic control.
- Growth failure due to erratic nutrient absorption.

#### Classic Celiac Disease – Features

- Three key components:
- 1. Villous atrophy
- 2. Symptoms of malabsorption (steatorrhea, weight loss, vitamin deficiency)
- 3. **Resolution** on gluten-free diet

# 😊 Latent Celiac Disease

- Diagnosed in **childhood**, recovered with gluten-free diet
- Remained silent even on gluten-containing diet
- CD can develop again later despite past normal biopsy

#### Potential Celiac Disease

- Normal biopsy
- Positive serology or genetics
- No treatment unless disease manifests
- Risk of developing CD in future → monitor only
- Important: Biopsy from distal duodenum (D2) is most involved site!

### Clinical Manifestations of Celiac Disease

### **GI and Nutritional:**

- Diarrhea with bulky, foul-smelling, floating stools
- Steatorrhea, flatulence, bloating
- Weight loss
- Growth retardation (children)
- Iron deficiency anemia
- Osteopenia, osteomalacia (Vit D/Ca def.)

#### Neurological:

- Fatigue, mood changes
- Ataxia, headache, epilepsy
- Depression, anxiety, idiopathic ataxia
- Peripheral neuropathy (Vit B12/Thiamine def.)

# Musculoskeletal:

- Arthritis (41% on gluten diet, 22% gluten-free)
  - Bone pain, pathological fractures

#### Hematologic:

- Iron def anemia (esp. in CD)
- Macrocytic anemia (B12)

#### Subclinical:

- Fatigue, mood changes
- Borderline iron deficiency
- Elevated liver enzymes (aminotransferases)

# Associated Conditions

| Condition                     | Notes  |
|-------------------------------|--|
| Dermatitis herpetiformis (DH) | Itchy papulovesicles; granular IgA deposits. Seen in 24% of CD, 85% of DH have CD. |
| T1DM                          | 2.6–7.8% of T1DM patients have CD  |
| Selective IgA Deficiency      | 2–3% → can cause false-negative serology   |

| Down, Turner, Williams syndromes                      | High association rates (4–19%) |
|---|--------------------------------|
| Autoimmune thyroiditis, liver disease, cardiomyopathy | Common in CD                   |

Celiac Disease increases risk for Enteropathy-associated T-cell lymphoma (EATL)

# Who Should Be Screened for CD?

#### GI Symptoms:

- Chronic diarrhea
- Weight loss, malabsorption
- Abdominal distention

#### Non-GI Symptoms/Findings:

- Short stature, delayed puberty
- Iron deficiency anemia
- Elevated aminotransferases
- Recurrent miscarriage or infertility

# High-Risk Groups:

- T1DM
- Autoimmune endocrinopathies
- First/second-degree relatives of CD patients
- Down, Turner, Williams syndromes
- Refractory anemia, IBS-like symptoms, ataxia

### Serologic Testing – Accuracy Table

| Antibody                        | Sensitivity | Specificity |
|---------------------------------|-------------|-------------|
| IgA anti-tTG (transglutaminase) | 90–98%      | 95–97%      |
| IgA EMA (endomysial Ab)         | 85–98%      | 97–100%     |
| IgA anti-gliadin Ab             | 80–90%      | 85–95%      |
| IgG anti-gliadin Ab             | 75–85%      | 75–90%      |

<sup>▲</sup> In IgA deficiency, use IgG-based tests

### Small Bowel Biopsy – Diagnosis

| Feature              | Description                          |
|----------------------|--------------------------------------|
| Mucosal inflammation | Lymphocytic infiltration             |
| Villous atrophy      | Flattened absorptive surface         |
| Crypt hyperplasia    | Elongated regenerative crypts        |
| Repeat biopsy        | After 3–6 months on gluten-free diet |

Biopsy = gold standard for diagnosis

### Management of Celiac Disease

# Core Principles:

- Patient education
- X Lifelong gluten-free diet
- Correct nutritional deficiencies (Fe, B12, Ca, Vit D)
- Access support groups
- Regular follow-up with a multidisciplinary team

# Ø Gluten-Free Diet − What to Avoid?

| X Gluten Sources |
|------------------|
| Wheat            |
| Barley           |
| Rye              |

Allow: corn, rice, potato, quinoa, oats (if uncontaminated)

# **CELIAC WHEAT – Disease Core**

- C Crypt hyperplasia (biopsy finding)
- E EMA / anti-TTG antibodies (best serologic tests)
- $L-\bigcirc$  Lymphocytic infiltration (immune cells in biopsy)
- $I \langle \rangle$  Iron deficiency anemia (microcytic type)
- $A \bigcap$  Atrophy of villi (flattened lining = hallmark)
- C Classic triad: Steatorrhea, Weight loss, Diet response
- $W \bigcirc$  Wheat (major gluten source)
- H # HLA-DQ2/DQ8 (genetic risk)
- **E** − **( Enteropathy** (chronic gut inflammation)
- $A \bigwedge$  Autoimmune links (like T1DM, thyroiditis)
- T O T-cell lymphoma (EATL risk, serious stuff!)

# **♣** BAD GAS From CELIAC – Symptoms

- B Q Bloating
- A ( Anemia
- D 🗐 Diarrhea
- G S Growth retardation (especially in kids)
- A Arthritis
- S **Steatorrhea** (floating, greasy poop)
- F 🚱 Fatigue
- C **⇔** Cheilitis/Glossitis (cracked lips, smooth tongue)
- E 4 Electrolyte abnormalities (Ca, Mg, K issues)
- L Liver enzymes elevated
- I **○ ②** Infertility
- **A** − **A Ataxia** (balance problems)
- $C \bigcirc \textcircled{A}$  Constipation sometimes (yep, not always diarrhea)

# **CANCER BAD – Complications**

- C Carcinoma of the small bowel
- $A \langle \rangle$  Anemia (chronic)
- $N \bigcirc$  Neuropathy (B12-related)
- C Crypt hyperplasia persists (active disease)
- E − ♠ EATL − T-cell lymphoma
- **R** − ☑ **Refractory CD** (won't respond to diet)
- B − **Sone fractures** (osteopenia/osteomalacia)
- A Amenorrhea
- **D** − **□ Dermatitis herpetiformis** (itchy rash)

# **A** TEG – Serologic Tests

- T − *Ø* Tissue transglutaminase (anti-TTG)
- E 
  Endomysial antibody (EMA)
- G Gliadin antibody (IgA or IgG)

 $\triangle$  If IgA deficient  $\rightarrow$  use IgG-based tests instead!

# **♦** G-FREE – Management

- G − **X ⑤** Gluten-free diet (forever and ever!)
- **F** − **♦ Fix deficiencies** (iron, B12, D, Ca)
- R 31 Regular follow-up (labs, bone health, weight)
- E − **②** Educate patient/family (label reading champs!)
- E Q Evaluate for complications (lymphoma, fractures, etc.)

# ✓ ABC – Classic Biopsy Findings in Celiac Disease

When a small bowel biopsy is done (usually from the second part of the duodenum − D2), these are the classic histological findings:

| Letter | Meaning           | Explanation  |
|--------|-------------------|--|
| A      | Atrophy of villi  | Flattened or absent villi = reduced surface area for absorption            |
| В      | Blunted mucosa    | Mucosal surface looks smooth and distorted                                 |
| C      | Crypt hyperplasia | Glandular structures (crypts) are elongated and increased in number due to |
|        |                   | mucosal regeneration efforts   |

# **Q** Bonus Tip for Exams:

If you see villous atrophy + crypt hyperplasia + increased intraepithelial lymphocytes, think CELIAC Biopsy is still the gold standard for diagnosis (especially when combined with positive serology).