INFLAMMATORY BOWEL DISEASE

Jean-Paul Achkar, MD
Center for Inflammatory Bowel Disease
Cleveland Clinic



WHAT IS INFLAMMATORY BOWEL DISEASE (IBD)?

Chronic inflammation of the intestinal tract

- Two related but different diseases:
 - Ulcerative colitis
 - Crohn's disease

WHAT IBD IS NOT

• IBD is sometimes confused with:

- Irritable bowel syndrome (IBS)
- Diverticulitis
- "Colitis"

INTESTINES: NORMAL STATE

• Protective immune cells are present in intestinal wall

• Immune system turns on and off to fight harmful substances like bacteria and viruses that pass through intestines

Microflora distribution in the GI tract

Oral cavity - 10^7 to 10^8

Stomach - 0 to 10³

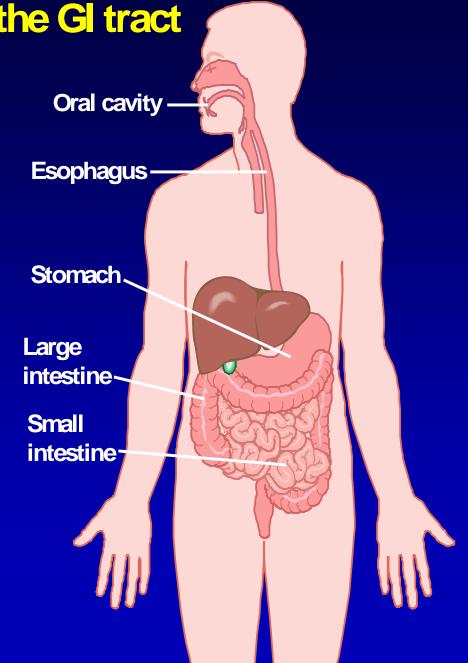
Duodenum - < 10^8

Jejunum - < 10³ to 10⁵

 $10^{5} \text{ to } 10^{7}$

Large intestine - 10¹⁰ to 10¹²

Stool - 10¹⁰ to 10¹²

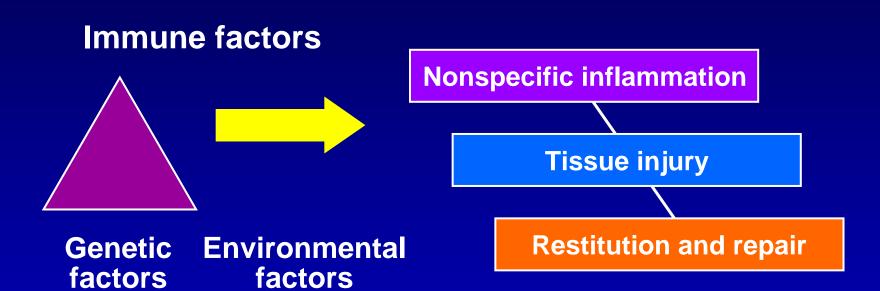


WHAT HAPPENS IN IBD?

• The immune system is activated by some unidentified factor

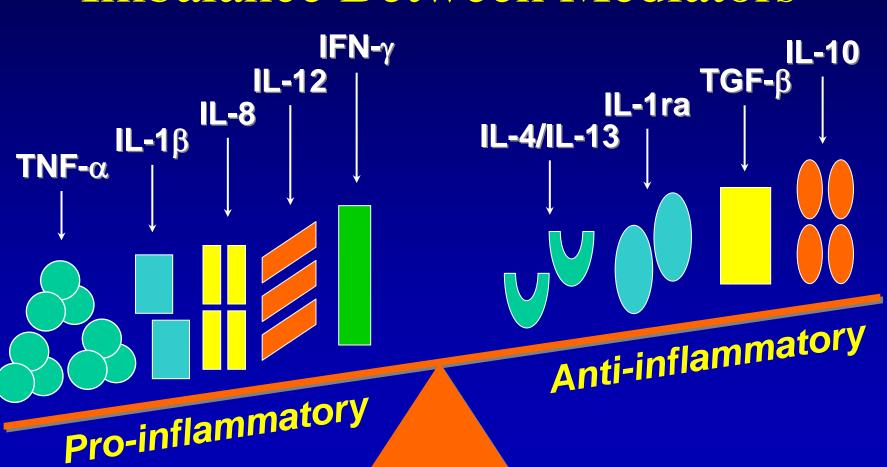
- The immune system does not turn off:
 - Uncontrolled inflammation
 - Attack on normal intestinal cells

Multifactorial Basis of IBD



Elson CO et al. In: *Proceedings of V International Symposium on Inflammatory Bowel Diseases*, 1998.

Chronic Inflammation: Imbalance Between Mediators



HOW COMMON IS IBD?

- More than 1 million cases estimated in the United States:
 - Ulcerative colitis: 50%
 - Crohn's disease: 50%
- New cases diagnosed at a rate of 10 cases per 100,000 people

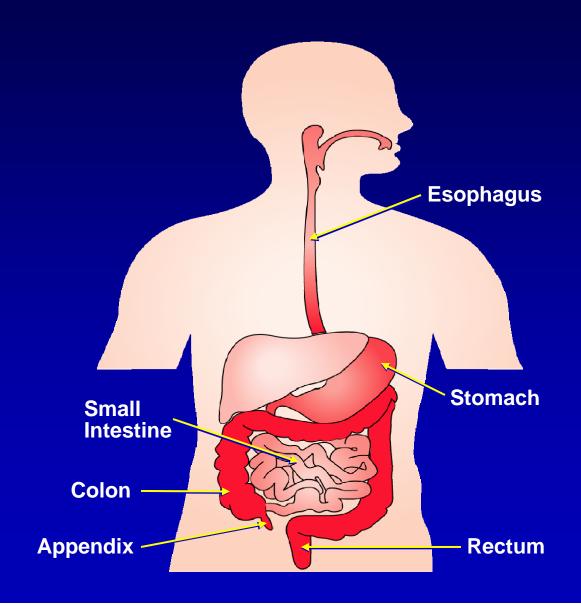
SCOPE OF THE PROBLEM

- Chronic lifelong diseases
- Periods of active disease alternating with periods of disease control:
 - Other conditions can cause symptoms that mimic those of IBD
- Complications can develop
- Surgery frequently required

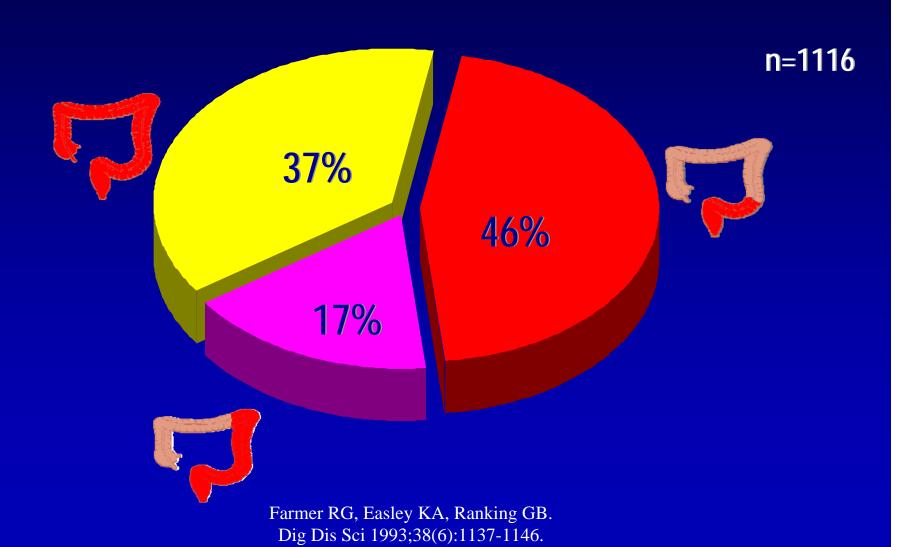
ULCERATIVE COLITIS

- Inflammation of the inner lining of the colon (mucosa)
- Only the colon is involved- starts in the rectum and spreads up the colon in continuous pattern
- Curable by surgery- removal of the colon

Gastrointestinal Tract



Disease Distribution at Presentation

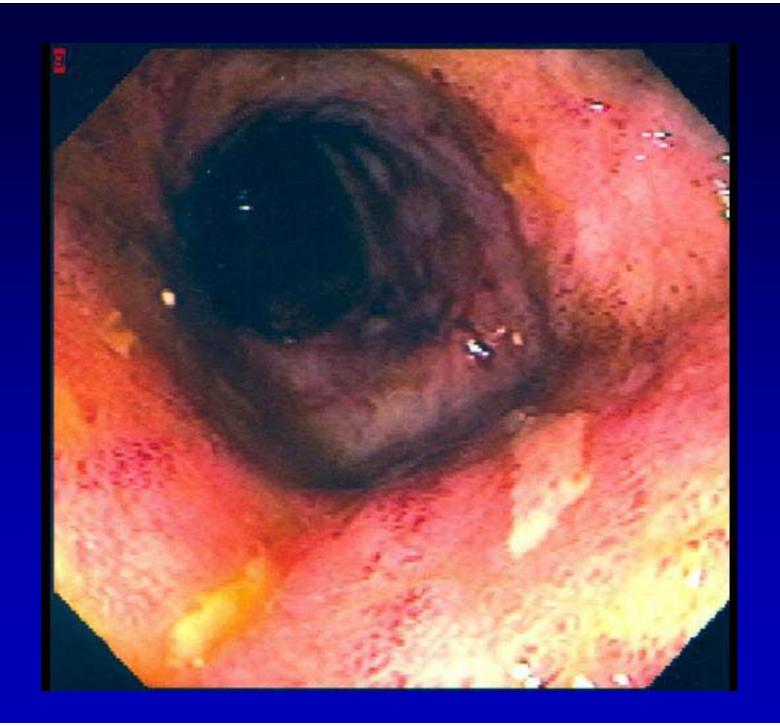


ULCERATIVE COLITIS: SYMPTOMS

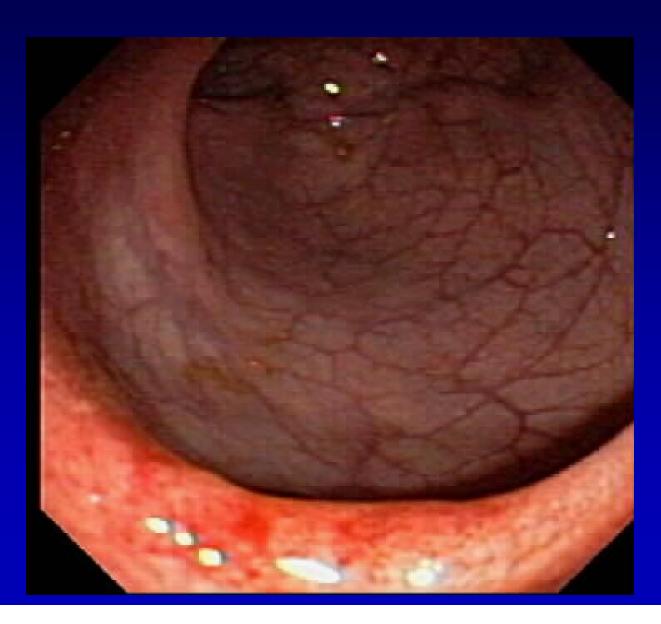
- Diarrhea
- Rectal bleeding
- Rectal urgency
- Abdominal cramps
- Fever

Normal Colonic Mucosa

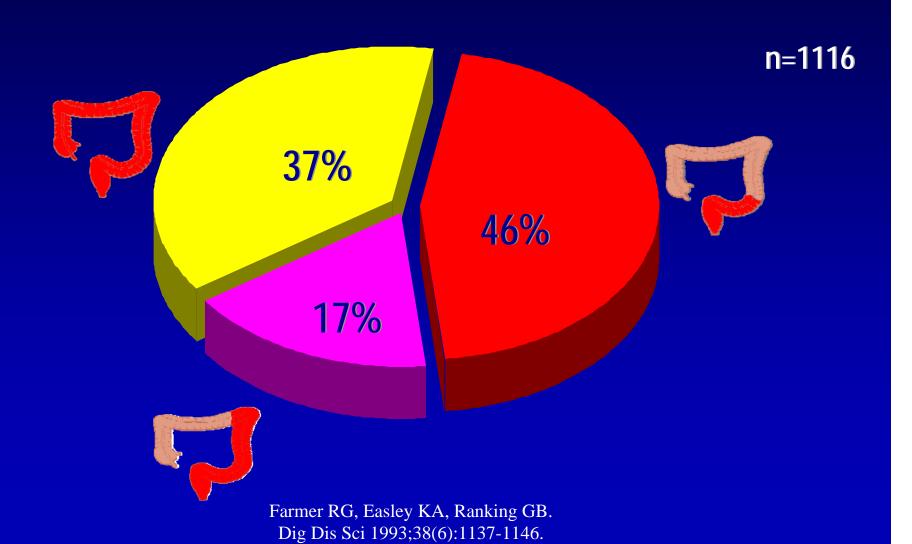




UC: Transition Point



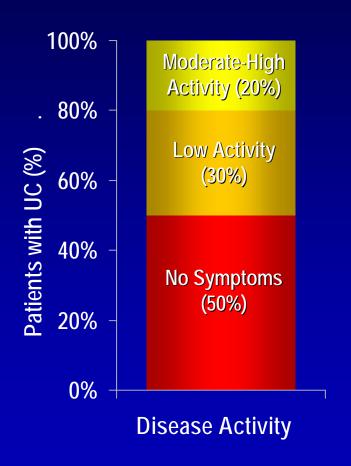
Disease Distribution at Presentation

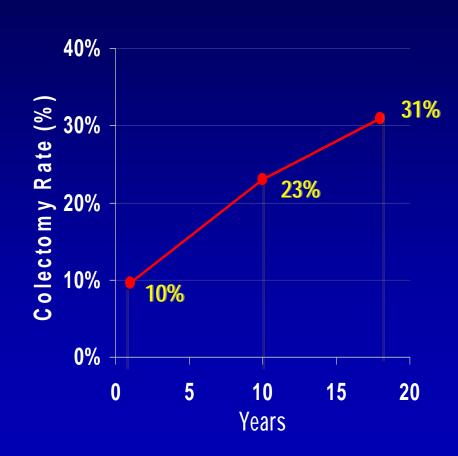


ULCERATIVE COLITIS: SYMPTOMS

- Diarrhea
- Rectal bleeding
- Rectal urgency
- Abdominal cramps
- Fever

UC Natural History

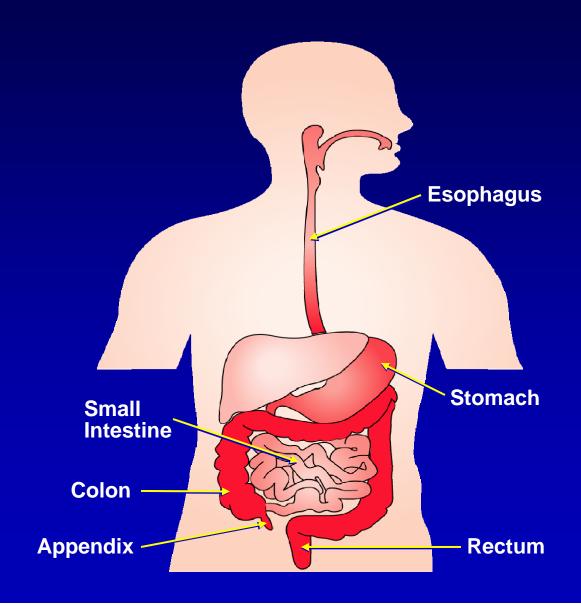




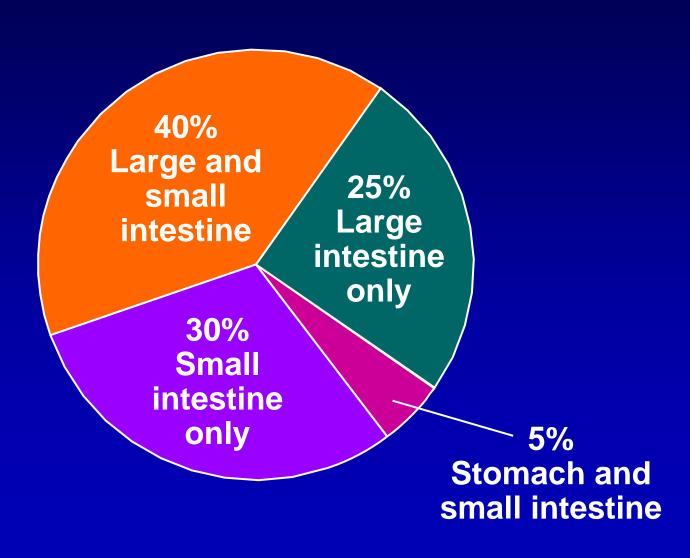
CROHN'S DISEASE

- Inflammation can involve all layers of the intestinal wall
- Can involve any part of the intestines from mouth to rectum with skip lesions- areas in between disease can be normal
- Usually comes back after surgery

Gastrointestinal Tract

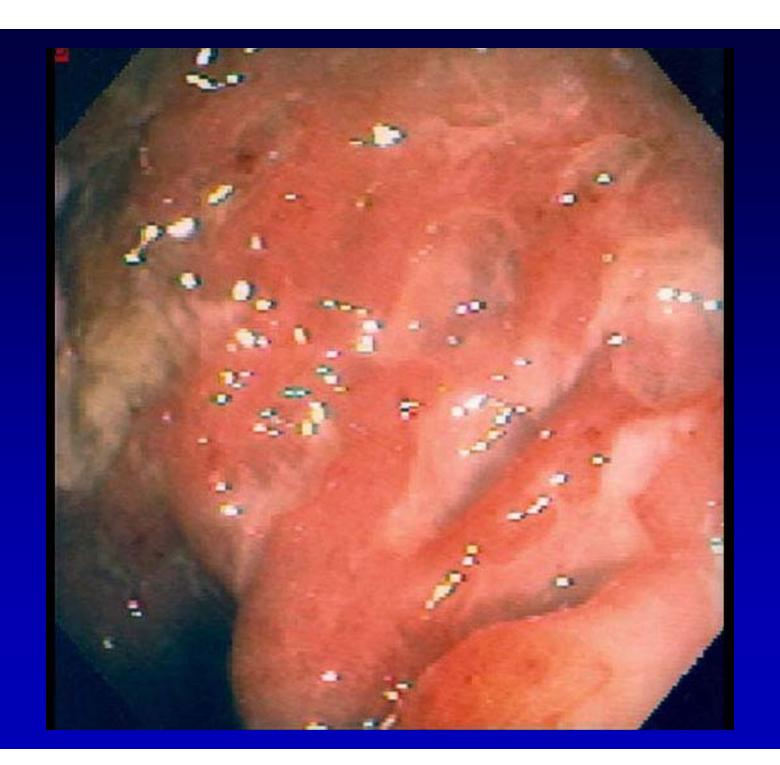


Locations in the GI Tract Most Often Affected



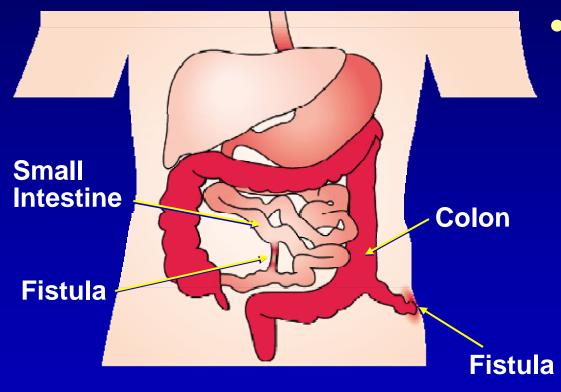
CROHN'S DISEASE: SYMPTOMS

- Same as those for ulcerative colitis
- Weight loss
- Fistulas- abnormal connection between intestine and other organs
- Abscess- collection of pus
- Strictures- areas of narrowing





Crohn's Disease Complications: Fistulas



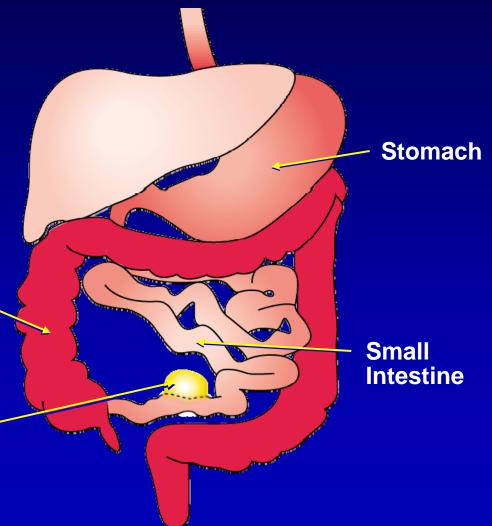
• A tunnel between two sections of the intestines or between the intestines and other organs, including the skin

Crohn's Disease Complications: Abscesses

• A localized collection of pus within the tissue of the GI tract

Large Intestine (Colon)

Abscess from a fissure in the small intestine into the peritoneal cavity



Complications of CD: Fistulas

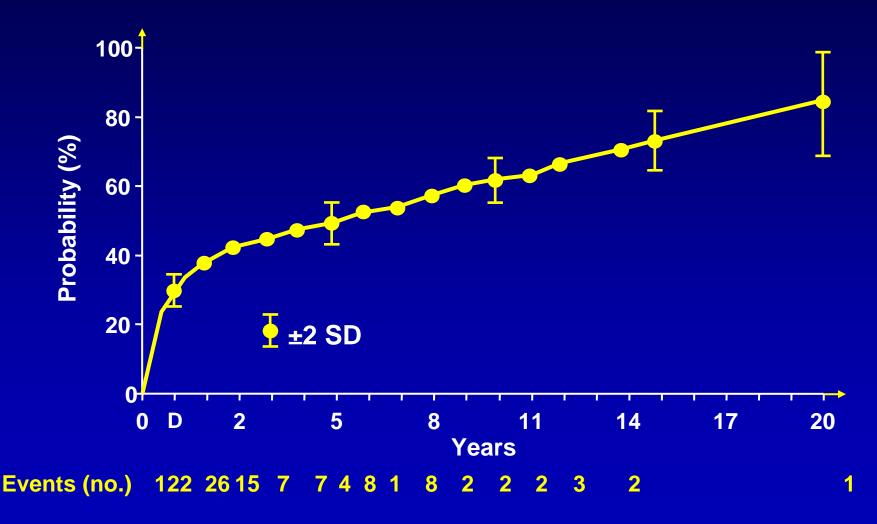
Abdominal Fistula



Perianal Fistula



Cumulative Probability of Surgical Intervention in CD



Munkholm P, et al. *Gastroenterology*. 1993;105:1716.

Extraintestinal Manifestations

- Skin disorders:
 - Erythema nodosum
 - Pyoderma gangrenosum
- Joint disorders:
 - Peripheral arthritis
 - Sacroiliitis
 - Ankylosing spondylitis
- Ocular disorders:
 - Iritis, uveitis, and episcleritis

Extraintestinal Manifestations

- Hepatobiliary:
 - Gallstones
 - Primary sclerosing cholangitis (PSC)
 - Cholangiocarcinoma
- Renal:
 - Renal stones
 - Amyloidosis
- Other manifestations:
 - Aphthous stomatitis
 - Hypercoagulable state

Episcleritis



Peripheral Arthritis



Erythema Nodosum



Pyoderma Gangrenosum



Treatment of IBD

Goals of Treatment

- Induce response/remission
- Maintain response/remission
- Heal mucosal lining
- Prevent or cure complications (eg, fistulas)
- Improve quality of life
- Restore and maintain nutrition
- Limit surgery

Treatment Options for IBD

- 5-ASA agents
- Antibiotics
- Steroids
- Immunomodulators:
 - 6-MP/azathioprine
 - Methotrexate
- Biologic agents:
 - Anti-TNF agents
 - Natalizumab