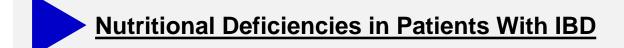
Nutrition in Patients With Inflammatory Bowel Disease (IBD)



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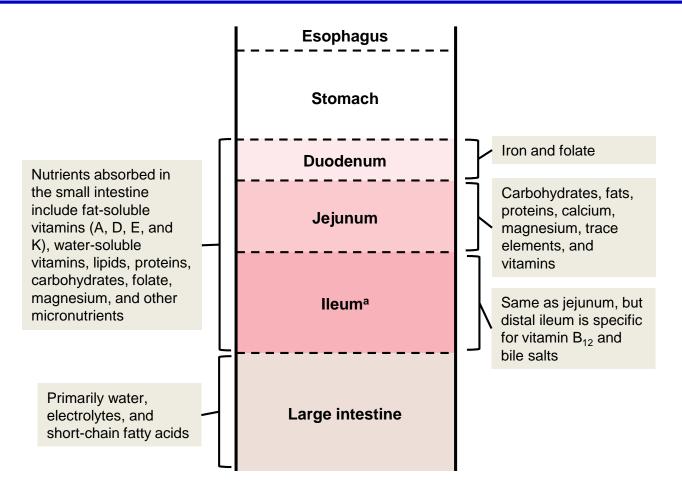
Diet and Nutrition in Patients With IBD

- Nutrition therapy is an important aspect of IBD disease management, in addition to pharmacological therapy or surgery^{1,2}
- Goals of nutrition therapy include the following¹⁻³:
 - Promote healing, immunity, and energy levels
 - Prevent or minimize symptoms of IBD

- Address malnutrition and ensure optimal growth
- Improve efficacy of medications
- Diet is the primary behavioral factor that can be manipulated by patients with IBD³
 - Patients who seek to improve their underlying condition may alter their diet through food avoidance or experiment with trial-anderror processes to identify foods that trigger symptoms⁴
- No diet has been shown to prevent IBD, but nutrition-related factors have been associated with a decreased risk of new-onset IBD¹
 - Breastfeeding is associated with a lower risk of childhood onset of IBD²⁻⁴
 - Diets rich in fruits and vegetables and low in animal fats and sugar are associated with a decreased risk of IBD¹
- Dietary factors may contribute to the development and pathogenesis of IBD by altering the microbiota, metabolome, host-barrier function, and innate immunity⁵
- Concerns regarding the side effects of IBD medications (eg, biologics) may prompt patients to seek dietary guidance^{1,6}



Nutrient Absorption in the GI Tract



- Nutrient absorption takes place in various sites of the GI tract¹
- Nutrient absorption may be compromised in patients with IBD, depending on the location and severity of intestinal inflammation^{2,3}
 - Small intestine inflammation can result in malabsorption of macro- and micronutrients²
 - Large intestine inflammation can result in malabsorption of water and electrolytes²

Figure modified from Gropper SS, et al.1

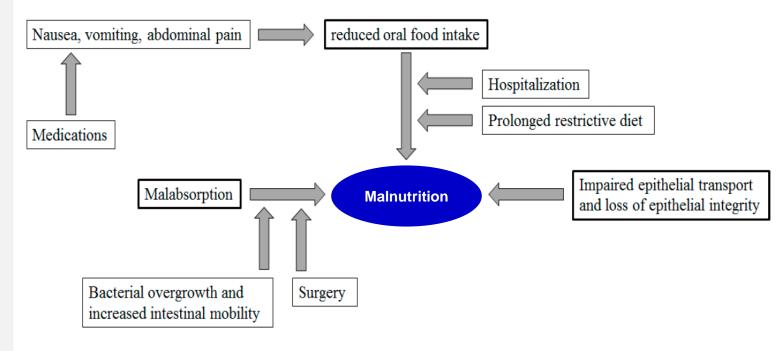
^{1.} Gropper SS, et al. Advanced Nutrition and Human Metabolism. 5th ed. Wadsworth/Cengage Learning; 2009. 2. Crohn's & Colitis Foundation. https://www.crohnscolitisfoundation.org/diet-and-nutrition/malnutrition-and-ibd. Accessed May 1, 2020. 3. Balestrieri P, et al. Nutrients. 2020;12(2):372.

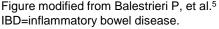


^aMany additional nutrients may be absorbed from the ileum depending on the transit time. GI=gastrointestinal; IBD=inflammatory bowel disease.

Factors Contributing to Malnutrition in Patients With IBD

- Malnutrition refers to deficiencies, excesses, or imbalances in a person's intake of energy or nutrients¹
- Many factors can contribute to nutritional inadequacies in patients with IBD:
 - Behavioral factors: alterations in diet through food avoidance or trial-and-error processes to identify foods that trigger symptoms²⁻⁴
 - Physical factors: malabsorption, small intestinal bacterial overgrowth, and surgery^{5,6}
 - Medication: IBD medications such as corticosteroids, sulfasalazine, and methotrexate⁷
 - Symptoms of IBD: diarrhea, nausea, vomiting, and abdominal pain^{5,7}





^{1.} World Health Organization. https://www.who.int/news-room/fact-sheets/detail/malnutrition. Accessed May 1, 2020. 2. Casanova MJ, et al. *J Crohns Colitis*. 2017;11(12):1430-1439. 3. Halmos EP, et al. *Nat Rev Gastroenterol Hepatol*. 2015;12(3):133-146. 4. Romano A, Castagna V. In: *Human Nutrition From the Gastroenterologist's Perspective*. Springer International Publishing Switzerland; 2016:79-98. 5. Balestrieri P, et al. *Nutrients*. 2020;12(2):372. 6. Wedrychowicz A, et al. *World J Gastroenterol*. 2016;22(3):1045-1066. 7. Crohn's & Colitis Foundation. https://www.crohnscolitisfoundation.org/diet-and-nutrition/malnutrition-and-ibd. Accessed May 1, 2020.



Prevalence of Malnutrition in Patients With IBD

- Reported prevalence of malnutrition in patients with IBD ranges between 20% and 85%¹
 - Malnutrition is a considerable problem in patients with CD, given its capacity to affect any part of the gastrointestinal tract—unlike UC, which is restricted to the colon^{2,3}
- The nutritional status of patients with IBD is frequently altered even when they are in remission. However, the severity of malnutrition in patients with IBD may also be related to disease activity, duration, and extent of disease/inflammation¹⁻⁵

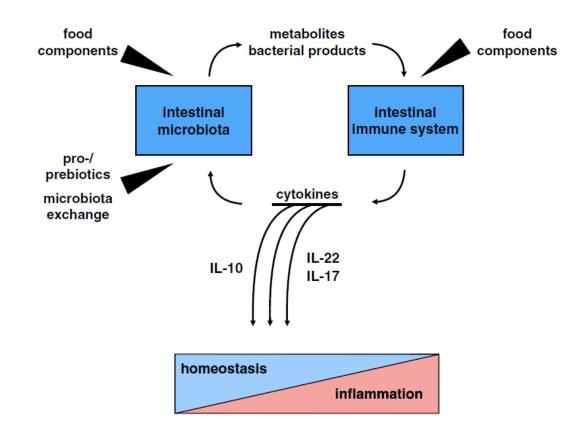
Malnutrition	CD	UC
Prevalence ⁴	• 65%-75%	• 18%-62%
Presentation ¹	May develop over a long period of time	May present during a severe acute flare
Inflammation and potential impact on nutritional deficiencies ¹⁻³	 Inflammation is patchy and may occur throughout the small and large bowel Ileal involvement may result in decreased nutrient absorption. Protein-energy and specific nutrient malnutrition is more common in patients with CD 	 Continuous and uniform inflammation confined to the colon Patients with UC may have less significant nutrient deficiencies, although severe diarrhea and blood loss can cause weight loss and anemia



Role of Diet, Microbiome, and Inflammation in IBD

- Intestinal microbiota dysbiosis is repeatedly seen in patients with IBD and is recognized as a key factor in gut inflammation¹
 - The dominant phyla in a normal gut are Firmicutes and Bacteroidetes, which account for approximately 64% and 23% of intestinal flora, respectively²
 - Intestinal microbiota can be beneficial and promote, for example, fermentation of indigestible carbohydrates, absorption of complex lipids, or vitamin synthesis²
 - Various studies have demonstrated a marked reduction in Firmicutes within the microbiota of patients with IBD²
 - Some bacteria, such as those from the phylum Proteobacteria, have been shown to have pathogenic potential in the development of IBD²
- A homeostatic balance of the host-bacteria relationship is important and vital for a normal health process¹

Interactions Between Diet, Microbiota, and Immune System in the Intestine²





Burden of Malnutrition in Patients With IBD

- Malnutrition has many detrimental effects¹
 - It is associated with deterioration in muscle, respiratory, and immune function
 - It may delay wound healing and recovery from illness
- Malnutrition is associated with poor outcomes in patients with IBD²
 - It is an independent risk factor for venous thromboembolism, nonelective surgery, and increased mortality in patients with IBD
 - It is also associated with a higher frequency of postoperative complications, longer hospital stays, decreased quality of life, and higher health costs

A Nationwide Study Using Hospital Discharge Data^a (1998-2004) in 52,142 Patients With IBD^b and Malnutrition^{1,c}

	In-hospital	Increase in	Increase in
	mortality,	length of stay,	hospital charges,
	OR (95% CI)	% (95% CI)	% (95% CI)
Malnutrition	3.49 (2.89-4.23)	55 (52-59)	57 (52-62)

- The adjusted OR for malnutrition among IBD admissions compared with non-IBD admissions was 5.57 (95% CI: 5.29-5.86)
- Hospital length of stay for IBD admissions with malnutrition was more than twice that of those without the diagnosis (11.9 days versus 5.8 days, *P*<0.00001)
- A greater than 2-fold difference in average hospital charges between those with and without malnutrition was observed (\$45,188 versus \$20,295, P<0.0001)

Note: The study was limited by the underlying disease severity in patients with IBD. Patients with severe disease may be more malnourished and may concurrently have a greater risk of in-hospital mortality.

^aData extracted from Nationwide Inpatient Samples, which is the largest all-payer database of national hospital discharges. ^bA total of 36,448 patients with CD and 15,694 patients with UC were included in the study. ^cClinically diagnosable protein-energy malnutrition.

CD=Crohn's disease; CI=confidence interval; IBD=inflammatory bowel disease; OR=odds ratio; UC=ulcerative colitis.

^{1.} Nguyen GC, et al. Inflamm Bowel Dis. 2008;14(8):1105-1111. 2. Bischoff SC, et al. Clin Nutr. 2020;39(3):632-653.

Clinical Aspects of Malnutrition in Patients With IBD



Types of Malnutrition in Patients With IBD

- Malnutrition comprises 3 broad groups of conditions¹:
 - Undernutrition: wasting (low weight for height), stunting (low height for age), and underweight (low weight for age)
 - Overnutrition: overweight, obesity, and diet-related noncommunicable diseases
 - Micronutrient-related malnutrition: lack of specific important vitamins and minerals or micronutrient excess
- Malnutrition is common among patients with IBD and includes conditions of all 3 categories²



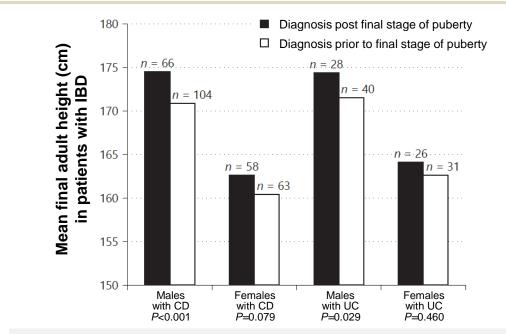
Protein-Energy Malnutrition in Patients With IBD

- Patients with IBD often do not consume an adequate number of calories¹
 - Decreased hunger, reduced sensation of pleasure related to eating, and changes in mood may contribute to reduced caloric intake²
 - Self-imposed elimination diets (in an attempt to control symptoms and flares or prevent disease relapse) may also result in malnutrition¹⁻³
- A prospective multicenter study of 333 patients with IBD in Spain showed that 76% of patients avoided some foods to prevent disease relapse, whereas 86% did so during active disease due to fear of worsening the flare²
 - Commonly excluded foods included fats, proteins, dairy products, and vegetables

Note: There could be a bias that most patients who agreed to participate in the nutritional assessment were those who had disease activity, even though consecutive patients in clinical remission or who had mild activity were included.

 In pediatric patients with IBD, malnutrition contributes to disrupted pubertal development and impaired growth velocity and can result in growth failure^{1,4,5}

Pediatric Patients Diagnosed With IBD Prior to Puberty May Experience Reduction in Growth⁵



Male gender, younger age at diagnosis, diagnosis prior to final stage of puberty, and lower height z-score at diagnosis impose risk for reduced adult height in both CD and UC

Note: This study was limited by its retrospective design and did not enable data collection on height velocity, bone age, and pubertal staging at diagnosis.



Altered Body Composition and Overnutrition in Patients With IBD

- Poor dietary intake, increased rates of protein turnover and loss of nutrients during active disease, or effects of IBD treatments
 contribute to alterations in body composition^{1,2}
 - Low muscle mass (sarcopenia), strength, and performance have been reported in adult patients with IBD²
 - A systematic review of patients with IBD (N=658 across 5 studies) reported that 42% of patients with IBD had sarcopenia³
 Note: This study was limited by its retrospective nature and the heterogenous definitions of sarcopenia, determined by 1 or more radiologic assessments of body mass composition across the studies.
 - Patients affected by sarcopenia may present with normal BMI or be overweight or obese (sarcopenic obesity) and therefore
 may not be identifiable as undernourished using traditional measures^{1,4-6}
 - Inadequate body composition is associated with poor outcomes such as postoperative complications, longer hospital stays, and increased healthcare costs⁵
- Patients with IBD may experience overnutrition and become overweight or obese (including during remission)^{1,2,6}
 - A study reviewing medical records of patients with IBD^a (from 2000 to 2012; N=581) reported obesity (BMI≥30) in 32.7% of adult patients with IBD (30.3% for CD and 35.2% for UC)⁷

Note: This study was limited by its retrospective nature and the use of BMI, which has a poor linear relationship with total body fat. Additionally, the lack of measurement for mesenteric fat, which may be more important to inflammation than subcutaneous fat, may limit the interpretability of the data.



Common Micronutrient Deficiencies in Patients With IBD

- Type of micronutrient and vitamin deficiency in patients with IBD depends on many factors, including disease localization, activity, and medication use for IBD¹
- Vitamin supplementation may correct most deficiencies but cannot guarantee adequacy; iron, zinc, and vitamin D are likely to require specific replacement regimens²

Micronutrient Deficiencies³

Micronutrient	At-risk individuals	Food sources	Common consequences of deficiency
Iron	Those with active disease; vegetarians and vegans; premenopausal women	Red meat; offal	Anemia, fatigue, weakness, brittle nails
Zinc	Vegetarians and vegans; chronic diarrhoea	Meat; fortified cereals	Impaired healing, disturbed smell and taste, delayed growth in children
Magnesium	Chronic or severe acute diarrhoea	Leafy-green vegetables; soybean	Disturbed bone health, muscular cramps, fatigue
Calcium	Restriction of dairy	Dairy; calcium-fortified dairy alternatives	Decreased bone density

Micronutrient	At-risk individuals	Food sources	Common consequences of deficiency
Vitamin B ₁₂	Vegetarians and vegans; ileal disease or resection	Animal-based foods	Anemia, fatigue, neurological effects
Vitamin D	Dark-skinned patients; those with decreased exposure to UV rays	Limited amount in fortified foods (e.g. margarine, milk)	Disturbed calcium homeostasis and bone health; possible enhancement of inflammatory activity
Folate	Those on restrictive or elimination diets; sulfasalazine therapy	Whole grains; leafy-green vegetables; fortified cereals	Anemia, fatigue



Conditions and Complications of Malnutrition in Patients With IBD

- Malnutrition can lead to complications in patients with IBD, including sarcopenia, osteopenia, obesity, and anemia¹
- In addition to micronutrient and vitamin loss, water loss can result in dehydration²

Condition	Prevalence and characteristics in patients with IBD	
Low bone mass, strength, and osteoporosis ¹	 Estimated prevalence: 20% to 50% Deficiencies of calcium, vitamins, and other micronutrients can lead to low bone mass and osteoporosis Other contributing factors include overall cumulative corticosteroid exposure, extensive small-bowel disease or resection, chronic inflammation, and lack of physical activity 	
Anemia ¹	 Estimated prevalence: 36% to 90% Iron deficiency is the main cause of anemia in patients with IBD 	
Sarcopenia ³	 Estimated prevalence: ~40% Contributing factors: poor dietary intake, increased rates of protein turnover and gut loss of nutrients during active disease 	
Obesity ^{4,5}	 Estimated prevalence: ~30% Contributing factor: overnutrition 	
Growth delays ⁶	 Approximately one-third of children with CD and one-tenth of children with UC have an adult height that is less than expected due to IBD Contributing factors: chronic inflammation and malnutrition, long-term corticosteroid use 	

Nutritional care in patients with IBD is intended to prevent malnutrition and micronutrient deficiencies¹



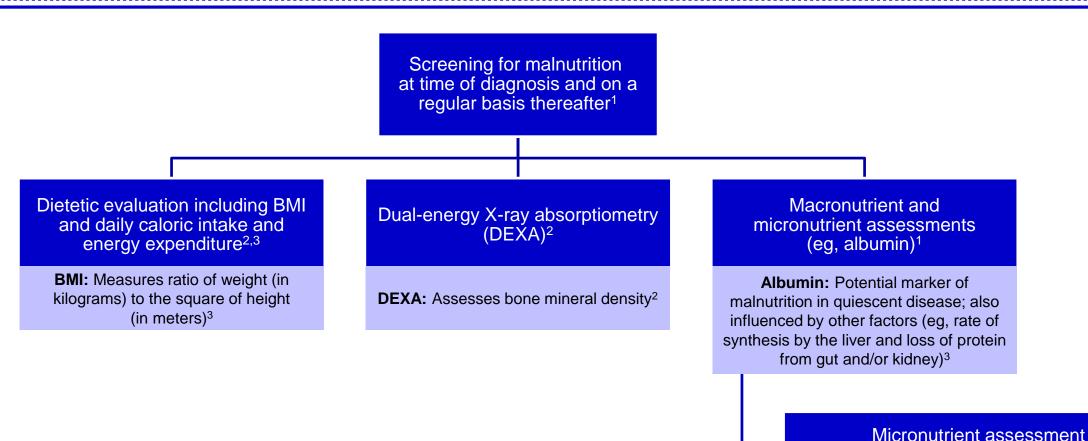
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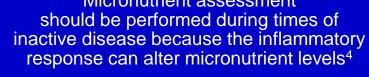
^{1.} Balestrieri P, et al. *Nutrients*. 2020;12(2):372. 2. Bischoff SC, et al. *Clin Nutr*. 2020;39(3):632-653. 3. Ryan E, et al. *Inflamm Bowel Dis*. 2019;25(1):67-73. 4. Halmos EP, Gibson PR. *Nat Rev Gastroenterol Hepatol*. 2015;12(3):133-146. 5. Flores A, et al. *Dig Dis Sci*. 2015;60(8):2436-2445. 6. Crohn's & Colitis Foundation. https://www.crohnscolitisfoundation.org/diet-and-nutrition/malnutrition-and-ibd. Accessed

Management of IBD With Nutrition



Assessment of Nutritional Status in Patients With IBD







Role of Registered Dietitian for Medical Nutrition Therapy

- It may be useful to partner with a registered dietitian to assist with nutritional interventions
- Dietitians may
 - Help tailor dietary modifications based on IBD history, anatomy, complications, comorbidities, and specific nutrient needs
 - Provide close monitoring through different phases of nutrition therapy (including advising on feasibility of or adherence to specific diets)
 - Address individualized meal planning
 - Answer patient questions and concerns that may arise during different stages of diet therapy

Dietary Approaches in IBD: Enteral Nutrition

- Enteral nutrition (EN): Use of a specific formula as nutritional therapy¹
 - EN is not currently a primary therapy option in UC but can be used for nutritional support²
 - Exclusive enteral nutrition (EEN): well-established method for inducing remission in children with recent-onset CD²
 - Clinical remission rates with EEN are 80% to 85% among children and adolescents with CD, and EEN has been shown to improve nutritional status, growth, and development^{2,3}
 - Evidence of efficacy in adults is significantly less than in children and likely related to practical issues (eg, disruption to normal life, poor palatability, lack of experience, lack of guidance) rather than mechanistic differences⁴
 - EEN is difficult to sustain as long-term maintenance therapy in CD³

A Retrospective Propensity Score–Matched Study^a Assessed the Impact of EN on Outcomes After Hospitalization in Patients With IBD (N=1578)^{5,b}

Patients ^b	30-Day hospital readmission, OR (95% CI)	30-Day mortality, OR (95% CI)
IBD	0.73 (0.52-1.02)	0.27 (0.08-0.90)
CD (n=800)	0.83 (0.54-1.28)	0.19 (0.04-0.99)
UC (n=778)	0.55 (0.32-0.97)	0.49 (0.08-3.09)

EN was associated with lower odds of readmission for UC and lower odds of mortality for CD

Note: This study was limited by its retrospective design, variation in EN provided, and the potential for bias introduced through variables chosen for propensity score matching.

^aNationwide Readmissions Database (2010-2015). ^bPatients with IBD and protein-energy malnutrition.

CD=Crohn's disease; CI=confidence interval; IBD=inflammatory bowel disease; OR=odds ratio; UC=ulcerative colitis.

^{1.} Crohn's & Colitis Foundation. https://www.crohnscolitisfoundation.org/diet-and-nutrition/nutritional-support-therapy. Accessed June 10, 2020. 2. Romano A, Castagna V. In: *Human Nutrition From the Gastroenterologist's Perspective*. Springer International Publishing Switzerland; 2016:79-98. 3. Green N, et al. *Nutrients*. 2019;11(5):947-960. 4. Ashton JJ, et al. *Clin Nutr*. 2019;38(1):80-89. 5. LeBrett WG, et al. Poster presented at: Crohn's & Colitis Congress; January 23-25, 2020; Austin, TX.

Findings for EN Therapy in Pediatric Patients With CD

- PEN and EEN diets as well as anti-TNF therapy have been shown to improve clinical symptoms of active CD¹
 - However, greater mucosal healing was observed with EEN or anti-TNF therapy compared with PEN
- Oral consumption of enteral formula may be as efficacious as continuous nasogastric feeding to induce remission and mucosal healing, with similar high compliance rates²
- The efficacy of EEN may be influenced by the location and the extent of intestinal inflammation³
 - Children with disease in the colon have been shown to respond better to EEN if the ileum is also involved (due to potential differences in underlying inflammatory mechanisms)



Dietary Approaches in IBD: Food-Based Treatment, Including Exclusion Diets

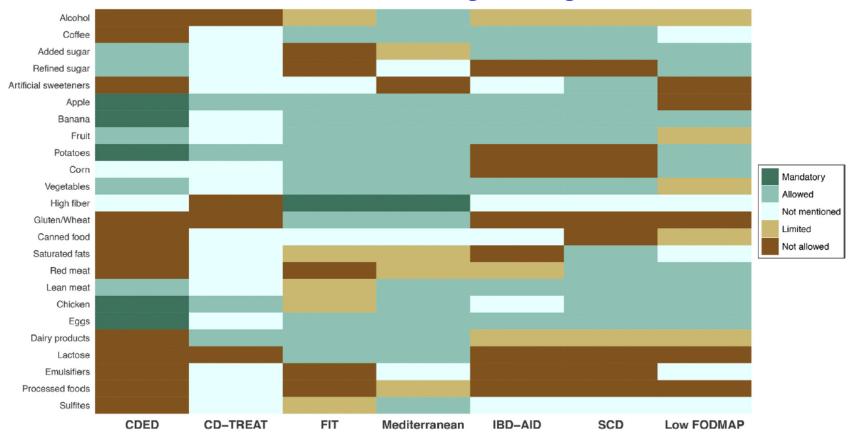
Appropriate patient selection for inclusion in trials of dietary therapy is critical, because the opportunity for early therapeutic escalation with medications that can prevent disease progression must

 Multiple diets are being investigated in clinical trials

be considered for nonresponders

- Food components that are recommended versus avoided in these trials vary widely
- None of the studies published to date suggest that dietary therapy should replace traditional therapy; however, these studies may elucidate how dietary intervention may be used in appropriately selected patients with adequate follow-up

Select Exclusion Diets Being Investigated in IBD





Select Investigational Nutritional Diets for the Management of IBD

- CD exclusion diet (CDED) with partial enteral nutrition (PEN): whole-foods diet for inducing sustained remission in patients with active CD^{1,2}
 - In a prospective study (n=78), CDED-PEN was as effective as EEN for inducing mucosal healing and sustained remission (75%) and was tolerated better than EEN by pediatric patients with mild to moderate CD¹
 Note: This study was limited by the indirect assessment of mucosal healing instead of endoscopy.
- CD treatment-with-eating diet (CD-TREAT): individualized food-based diet with similar composition to EEN³
 - In an RCT (n=25), CD-TREAT had similar effects to those of EEN on the gut microbiome and metabolome in healthy adults
 - In an open-label study (n=5), CD-TREAT led to a remission rate of 60% and decreased colonic inflammation in pediatric patients with active CD

Note: This study was limited by the small number of patients enrolled.

- Specific carbohydrate diet (SCD): diet that restricts complex carbohydrates and eliminates refined sugar^{4,5}
 - SCD is used in many different medical conditions, including IBD, irritable bowel syndrome, celiac disease, and autism⁴
 - In 2 prospective studies (n=10 and n=12), SCD improved mucosal healing and clinical and laboratory parameters and was
 associated with changes in gut microbial composition in pediatric patients with IBD^{4,5}

Note: The open-label study designs, small numbers of patients enrolled, and lack of control groups were the main limitations of these studies.



Dietary Guidance From the International Organization for the Study of IBD (IOIBD)

- The IOIBD recently provided expert opinion on specific dietary components, food groups, and food additives that may be prudent to increase or decrease in the diet to control and prevent relapse of IBD
 - Recommendations specific to patients with CD and UC are included
 - Guidance is based on the best current evidence available
 - The recommendations are not meant to exclude the role of nutritional assessment for malnutrition and correction of deficiencies when needed
 - For patients with persistent symptoms despite resolution of inflammation and absence of strictures, the IOIBD suggested that a low-FODMAP or lactose-free diet may improve symptoms

Dietary Guidance for Patients With CD and UC

Se		
nt to increa	CD recommendations	UC recommendations
Prudent to increase foods containing	Vegetables Fruits	Omega 3 oils from fish and food
Prudent to decrease foods containing	 Saturated and trans fat Emulsifiers Carrageenans Artificial sweeteners Maltodextrins 	 Red meat, processed meats Dairy fat, palm and coconut oil Saturated and trans fat Emulsifiers Carrageenans
	Titanium dioxide	Artificial sweetenersMaltodextrinsTitanium dioxide



Summary

- Dietary factors may contribute to the development and pathogenesis of IBD by altering the microbiota, metabolome, host-barrier function, and innate immunity
- Malnutrition affects 20% to 85% of patients with IBD and includes conditions such as proteinenergy malnutrition, altered body composition and overnutrition, and micronutrient deficiency
- Malnutrition can lead to complications including anemia, osteoporosis, sarcopenia, obesity, and growth delays (in children), and it is associated with decreased quality of life, poor treatment outcomes, and higher health costs
- Patients with IBD are encouraged to screen for malnutrition and partner with a registered dietitian for subsequent nutritional interventions
- Common dietary approaches under investigation to support disease management for patients with IBD include enteral nutrition, CDED, CD-TREAT, and SCD



Available Resources

- Crohn's & Colitis Foundation
 - Malnutrition and IBD
 - Nutritional support therapy
- World Health Organization
 - Malnutrition fact sheet and Q&A
- International Organization for the Study of Inflammatory Bowel Disease
- Malnutrition Advisory Group, a standing committee of the British Association for Parenteral and Enteral Nutrition
 - Online Malnutrition Universal Screening Tool (MUST)
- Pt-Global Platform
 - Online Patient-Generated Subjective Global Assessment (PG-SGA) questionnaire
- Nutrition4Kids Foundation
 - Nutrition4IBD.com



References

- Adams DW, Gurwara S, Silver HJ, et al. Sarcopenia is common in overweight patients with inflammatory bowel disease and may predict need for surgery. *Inflamm Bowel Dis*. 2017;23(7):1182-1186.
- Afzal NA, Davies S, Paintin M, et al. Colonic Crohn's disease in children does not respond well to treatment with enteral nutrition if the ileum is not involved. Dig Dis Sci. 2005;50(8):1471-1475.
- Ashton JJ, Gavin J, Beattie RM. Exclusive enteral nutrition in Crohn's disease: evidence and practicalities. Clin Nutr. 2019;38(1):80-89.
- Balestrieri P, Ribolsi M, Guarino MPL, et al. Nutritional aspects in inflammatory bowel diseases. Nutrients. 2020;12(2):372.
- Bischoff SC, Escher J, Hebuterne X, et al. ESPEN practical guideline: clinical nutrition in inflammatory bowel disease. Clin Nutr. 2020;39(3):632-653.
- British Association for Parenteral and Enteral Nutrition. Malnutrition universal screening tool. https://www.bapen.org.uk/pdfs/must/must_full.pdf. Accessed June 10, 2020.
- Casanova MJ, Chaparro M, Molina B, et al. Prevalence of malnutrition and nutritional characteristics of patients with inflammatory bowel disease. J Crohns Colitis. 2017;11(12):1430-1439.
- Cohen SA, Gold BD, Oliva S, et al. Clinical and mucosal improvement with specific carbohydrate diet in pediatric Crohn disease. *J Pediatr Gastroenterol Nutr.* 2014;59(4):516-521.
- Crohn's & Colitis Foundation. Malnutrition and IBD. https://www.crohnscolitisfoundation.org/diet-and-nutrition/malnutrition-and-ibd. Accessed May 1, 2020.
- Crohn's & Colitis Foundation. Nutritional support therapy. https://www.crohnscolitisfoundation.org/diet-and-nutrition/nutritional-support-therapy. Accessed June 10, 2020.
- Crohn's & Colitis Foundation. Navigating inflammatory bowel diseases using a nutrition care pathway.
 https://www.crohnscolitisfoundation.org/sites/default/files/legacy/assets/pdfs/esa/nutrition-slides-fnce-sympo.pdf. Accessed June 10, 2020.
- de Castro MM, Corona LP, Pascoal LB, et al. Impaired nutritional status in outpatients in remission or with active Crohn's disease classified by objective endoscopic and imaging assessments. Clin Nutr ESPEN. 2019;33:60-65.
- Flores A, Burstein E, Cipher DJ, et al. Obesity in inflammatory bowel disease: a marker of less severe disease. Dig Dis Sci. 2015;60(8):2436-2445.
- Green N, Miller T, Suskind D, et al. A review of dietary therapy for IBD and a vision for the future. *Nutrients*. 2019;11(5):947-950.
- Gropper SS, Smith JL, Groff JL. Advanced Nutrition and Human Metabolism. 5th ed. Wadsworth/Cengage Learning; 2009.



References

- Halmos EP, Gibson PR. Dietary management of IBD—insights and advice. Nat Rev Gastroenterol Hepatol. 2015;12(3):133-146.
- LeBrett WG, Sauk J, Berkeley N. Enteral nutrition therapy is associated with fewer readmissions and deaths among malnourished inpatients with inflammatory bowel disease. Poster presented at: Crohn's & Colitis Congress; January 23-25, 2020; Austin, TX.
- Lee D, Baldassano RN, Otley AR, et al. Comparative effectiveness of nutritional and biological therapy in North American children with active Crohn's disease. *Inflamm Bowel Dis.* 2015;21(8):1786-1793.
- Levine A, Sigall-Boneh R, Wine E. Evolving role of diet in the pathogenesis and treatment of inflammatory bowel diseases. *Gut.* 2018;67(9):1726-1738.
- Levine A, Wine E, Assa A, et al. Crohn's disease exclusion diet plus partial enteral nutrition induces sustained remission in a randomized controlled trial. *Gastroenterology*. 2019;157(2):440-450.
- Levine A, Rhodes JM, Lindsay JO, et al. Dietary guidance from the International Organization for the Study of Inflammatory Bowel Diseases. Clin Gastroenterol Hepatol. 2020;18(6):1381-1392.
- Limdi JK, Aggarwal D, McLaughlin JT. Dietary practices and beliefs in patients with inflammatory bowel disease. *Inflamm Bowel Dis.* 2016;22(1):164-170.
- Martinez B, Dailey F, Almario CV, et al. Patient understanding of the risks and benefits of biologic therapies in inflammatory bowel disease: insights from a large-scale analysis of social media platforms. *Inflamm Bowel Dis.* 2017;23(7):1057-1064.
- Miele E, Shamir R, Aloi M, et al. Nutrition in pediatric inflammatory bowel disease: a position paper on behalf of the Porto Inflammatory Bowel Disease Group of the European Society of Pediatric Gastroenterology, Hepatology and Nutrition. *J Pediatr Gastroenterol Nutr.* 2018;66(4):687-708.
- Nazarenkov N, Seeger K, Beeken L, et al. Implementing dietary modifications and assessing nutritional adequacy of diets for inflammatory bowel disease. Gastroenterol Hepatol. 2019;15(3):133-144.
- Nguyen GC, Munsell M, Harris ML. Nationwide prevalence and prognostic significance of clinically diagnosable protein-calorie malnutrition in hospitalized inflammatory bowel disease patients. *Inflamm Bowel Dis.* 2008;14(8):1105-1111.
- Pigneur B, Ruemmele FM. Nutritional interventions for the treatment of IBD: current evidence and controversies. Ther Adv Gastroenterol. 2019;12:1756284819890534.
- Pt-Global. PG-SGA[©]. http://pt-global.org/?page_id=13. Accessed June 10, 2020.



References

- Rinawi F, Assa A, Almagor T, et al. Prevalence and predictors of growth impairment and short stature in pediatric-onset inflammatory bowel disease. Digestion. 2019; doi:10.1159/000501924.
- Romano A, Castagna V. Diet as therapy for IBD? In: Human Nutrition From the Gastroenterologist's Perspective. Springer International Publishing Switzerland; 2016:79-98.
- Rubio A, Pigneur H, Garnier-Lengline C, et al. The efficacy of exclusive nutritional therapy in paediatric Crohn's disease, comparing fractionated oral vs. continuous enteral
 feeding. Aliment Pharmacol Ther. 2011;33(12):1332-1339.
- Ryan E, McNicholas D, Creavin B, et al. Sarcopenia and inflammatory bowel disease: a systematic review. Inflamm Bowel Dis. 2019;25(1):67-73.
- Sabino J, Lewis JD. Treating inflammatory bowel disease with diet: a taste test. Gastroenterology. 2019;157(2):295-297.
- Scaldaferri F, Pizzoferrato M, Lopetuso LR, et al. Nutrition and IBD: malnutrition and/or sarcopenia? A practical guide. Gastroenterol Res Pract. 2017;2017:8646495.
- Sigall-Boneh R, Pfeffer-Gik T, Segal I, et al. Partial enteral nutrition with a Crohn's disease exclusion diet is effective for induction of remission in children and young adults with Crohn's disease. *Inflamm Bowel Dis.* 2014;20(8):1353-1360.
- Suskind DL, Cohen SA, Brittnacher MJ, et al. Clinical and fecal microbial changes with diet therapy in active inflammatory bowel disease. Clin Gastroenterol. 2018;52(2):155-163.
- Svolos V, Hansen R, Nichols B, et al. Treatment of active Crohn's disease with an ordinary food-based diet that replicates exclusive enteral nutrition. Gastroenterology. 2019;156(5):1354-1367.
- Wall CL, Day AS, Gearry RB. Use of exclusive enteral nutrition in adults with Crohn's disease: a review. World J Gastroenterol. 2013;19(43):7652-7660.
- Wedrychowicz A, Zając A, Tomasik P. Advances in nutritional therapy in inflammatory bowel diseases: review. World J Gastroenterol. 2016;22(3):1045-1066.
- World Health Organization. Fact sheets: malnutrition. https://www.who.int/news-room/fact-sheets/detail/malnutrition. Accessed May 1, 2020.
- World Health Organization. Q&A: malnutrition. https://www.who.int/features/ga/malnutrition/en/. Accessed May 1, 2020.
- Witkowski M, Witkowski M, Gagliani N, Huber S. Recipe for IBD: can we use food to control inflammatory bowel disease? Semin Immunopathol. 2018;40(2):145-156.

