Celiac disease

Celiac disease is a condition in which the immune system is abnormally sensitive to gluten, a protein found in wheat, rye, and barley. Celiac disease is an autoimmune disorder; autoimmune disorders occur when the immune system malfunctions and attacks the body's own tissues and organs. Without a strict, lifelong gluten-free diet, inflammation resulting from immune system overactivity may cause a wide variety of signs and symptoms involving many parts of the body.

Celiac disease can develop at any age after an individual starts eating foods containing gluten. The classic symptoms of the condition result from inflammation affecting the gastrointestinal tract. This inflammation damages the villi, which are small, finger-like projections that line the small intestine and provide a greatly increased surface area to absorb nutrients. In celiac disease, the villi become shortened and eventually flatten out. Intestinal damage causes diarrhea and poor absorption of nutrients, which may lead to weight loss. Abdominal pain, swelling (distention), and food intolerances are common in celiac disease. Inflammation associated with celiac disease may lead to an increased risk of developing certain gastrointestinal cancers such as cancers of the small intestine or esophagus.

Inflammation and poor nutrient absorption may lead to problems affecting many other organs and systems of the body in affected individuals. These health problems may include iron deficiency that results in a low number of red blood cells (anemia), vitamin deficiencies, low bone mineral density (osteoporosis), itchy skin rashes (dermatitis herpetiformis), defects in the enamel of the teeth, chronic fatigue, joint pain, poor growth, delayed puberty, infertility, or repeated miscarriages. Neurological problems have also been associated with celiac disease; these include migraine headaches, depression, attention-deficit/hyperactivity disorder (ADHD), and recurrent seizures (epilepsy). Many people with celiac disease have one or more of these varied health problems but do not have gastrointestinal symptoms. This form of the condition is called nonclassic celiac disease. Researchers now believe that nonclassic celiac disease is actually more common than the classic form.

Celiac disease often goes undiagnosed because many of its signs and symptoms are nonspecific, which means they may occur in many disorders. Most people who have one or more of these nonspecific health problems do not have celiac disease. On average, a diagnosis of celiac disease is not made until 6 to 10 years after symptoms begin.

Some people have silent celiac disease, in which they have no symptoms of the disorder. However, people with silent celiac disease do have immune proteins in their blood (antibodies) that are common in celiac disease. They also have inflammatory damage to their small intestine that can be detected with a biopsy.
In a small number of cases, celiac disease does not improve with a gluten-free diet and progresses to a condition called refractory sprue. Refractory sprue is characterized by chronic inflammation of the gastrointestinal tract, poor absorption of nutrients, and an increased risk of developing a type of cancer of the immune cells called T-cell lymphoma.

Frequency
Celiac disease is a common disorder. Its prevalence has been estimated at about 1 in 100 people worldwide.

Causes
The risk of developing celiac disease is increased by certain variants of the *HLA-DQA1* and *HLA-DQB1* genes. These genes provide instructions for making proteins that play a critical role in the immune system. The *HLA-DQA1* and *HLA-DQB1* genes belong to a family of genes called the human leukocyte antigen (HLA) complex. The HLA complex helps the immune system distinguish the body's own proteins from proteins made by foreign invaders such as viruses and bacteria.

The proteins produced from the *HLA-DQA1* and *HLA-DQB1* genes attach (bind) to each other to form a functional protein complex called an antigen-binding DQαβ heterodimer. This complex, which is present on the surface of certain immune system cells, attaches to protein fragments (peptides) outside the cell. If the immune system recognizes the peptides as foreign (such as viral or bacterial peptides), it triggers a response to attack the invading viruses or bacteria.

Celiac disease is associated with an inappropriate immune response to a segment of the gluten protein called gliadin. This inappropriate activation of the immune system causes inflammation that damages the body's organs and tissues and leads to the signs and symptoms of celiac disease.

Almost all people with celiac disease have specific variants of the *HLA-DQA1* and *HLA-DQB1* genes, which seem to increase the risk of an inappropriate immune response to gliadin. However, these variants are also found in 30 percent of the general population, and only 3 percent of individuals with the gene variants develop celiac disease.

It appears likely that other contributors, such as environmental factors and changes in other genes, also influence the development of this complex disorder.

Inheritance Pattern
Celiac disease tends to cluster in families. Parents, siblings, or children (first-degree relatives) of people with celiac disease have between a 4 and 15 percent chance of developing the disorder. However, the inheritance pattern is unknown.
Other Names for This Condition

• celiac sprue
• gluten enteropathy
• sprue

Diagnosis & Management

Genetic Testing Information

• What is genetic testing?
  /primer/testing/genetictesting

• Genetic Testing Registry: Celiac disease

Research Studies from ClinicalTrials.gov

• ClinicalTrials.gov
  https://clinicaltrials.gov/ct2/results?cond=%22celiac+disease%22

Other Diagnosis and Management Resources

• Beth Israel Deaconess Medical Center: Celiac Center

• GeneReview: Celiac Disease
  https://www.ncbi.nlm.nih.gov/books/NBK1727

• Massachusetts General Hospital Center for Celiac Research and Treatment
  https://www.massgeneral.org/children/services/treatmentprograms.aspx?id=1723

• MedlinePlus Encyclopedia: Celiac Disease Nutritional Considerations
  https://medlineplus.gov/ency/article/002443.htm

• North American Society for Pediatric Gastroenterology, Hepatology, and Nutrition: Gluten-Free Diet Guide

• University of Chicago Celiac Disease Center
  http://www.cureceliacdisease.org/?utm_source=celiacdisease&utm_medium=redirect&utm_campaign=CeliacdiseaseBRedirect
Additional Information & Resources

Health Information from MedlinePlus

- Encyclopedia: Celiac Disease
  https://medlineplus.gov/ency/article/000233.htm
- Encyclopedia: Celiac Disease Nutritional Considerations
  https://medlineplus.gov/ency/article/002443.htm
- Health Topic: Celiac Disease
  https://medlineplus.gov/celiacdisease.html

Genetic and Rare Diseases Information Center

- Celiac disease
  https://rarediseases.info.nih.gov/diseases/11998/celiac-disease

Additional NIH Resources

- National Institute of Diabetes and Digestive and Kidney Diseases: Celiac Disease
  https://www.niddk.nih.gov/health-information/digestive-diseases/celiac-disease
- National Institute of Diabetes and Digestive and Kidney Diseases: Dermatitis Herpetiformis, Skin Manifestations of Celiac Disease
  https://www.niddk.nih.gov/health-information/digestive-diseases/dermatitis-herpetiformis

Educational Resources

- Academy of Nutrition and Dietetics
  https://www.eatright.org/health/diseases-and-conditions/celiac-disease/celiac-disease-an-introduction
- KidsHealth: Celiac Disease
- MalaCards: celiac disease 1
  https://www.malacards.org/card/celiac_disease_1
- Orphanet: NON RARE IN EUROPE: Celiac disease
  https://www.orpha.net/consor/cgi-bin/OC_Exp.php?Lng=EN&Expert=555
- TeensHealth: Celiac Disease

Patient Support and Advocacy Resources

- Canadian Celiac Association
  https://www.celiac.ca/
- Celiac Disease Foundation
  https://celiac.org/
• Coeliac Australia
  https://www.coeliac.org.au/

• Gluten Intolerance Group
  https://gluten.org/

Clinical Information from GeneReviews
• Celiac Disease
  https://www.ncbi.nlm.nih.gov/books/NBK1727

Scientific Articles on PubMed
• PubMed
  https://www.ncbi.nlm.nih.gov/pubmed?term=%28Celiac+Disease%5BMAJR%5D%29+AND+%28celiac+disease%5BTI%5D%29+AND+review%5Bpt%5D+AND+english%5Bla%5D+AND+human%5Bmh%5D+AND+%22last+720+days%22%5Bdp%5D

Catalog of Genes and Diseases from OMIM
• CELIAC DISEASE, SUSCEPTIBILITY TO, 1
  http://omim.org/entry/212750

Medical Genetics Database from MedGen
• Celiac disease

Sources for This Summary


Citation on PubMed: https://www.ncbi.nlm.nih.gov/pubmed/20040864 
Free article on PubMed Central: https://www.ncbi.nlm.nih.gov/pmc/articles/PMC2830645/

Citation on PubMed: https://www.ncbi.nlm.nih.gov/pubmed/20212505

Citation on PubMed: https://www.ncbi.nlm.nih.gov/pubmed/20301720

Citation on PubMed: https://www.ncbi.nlm.nih.gov/pubmed/20661732 
Free article on PubMed Central: https://www.ncbi.nlm.nih.gov/pmc/articles/PMC2944025/

Citation on PubMed: https://www.ncbi.nlm.nih.gov/pubmed/21317918 
Free article on PubMed Central: https://www.ncbi.nlm.nih.gov/pmc/articles/PMC4003132/

Reprinted from Genetics Home Reference: 

Reviewed: April 2019 
Published: July 9, 2019

Lister Hill National Center for Biomedical Communications 
U.S. National Library of Medicine 
National Institutes of Health 
Department of Health & Human Services