

# Identifying Crohn's disease risk factors in the Ashkenazi Jewish population

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Credit: Susanna M. Hamilton, Broad Communications

It is estimated that one in three individuals of Ashkenazi Jewish (AJ) descent carry mutations that increase their risk for certain genetic diseases. For instance, Crohn's, a highly heritable inflammatory bowel

disease, is two to four times more prevalent among people of AJ ancestry, compared to those of non-Jewish European ancestry.

In a study published today in *PLoS Genetics*, an international team of researchers describes Crohn's disease-related genetic [risk factors](#) that appear up to four-fold times more frequently in the AJ population, compared to other populations.

Analysing 5,685 exomes (protein-coding regions of the genome) from individuals of AJ descent, the researchers provide a detailed list of genetic variants that increase the risks of Crohn's, as well as certain rare diseases like Gaucher, Canavan, and Tay Sachs. While Crohn's is an inflammatory disease of the digestive tract, Gaucher is a metabolic disease, and Canavan and Tay Sachs are both [neurodegenerative diseases](#).

"We wanted to make this information immediately available to the community. This will benefit other researchers who are studying these diseases and can help clinical practitioners with screening efforts," said lead author Manuel Rivas, assistant professor in the Department of Biomedical Data Science at Stanford University and visiting scientist at the Broad Institute of MIT and Harvard.

The researchers found ten Crohn's-associated mutations in the genes NOD2 and LRRK2, all of which occurred more frequently in the AJ population. NOD2 is an immune-system function gene, while LRRK2, which is active in the brain, is associated with Parkinson's [disease](#).

They also noted eight-, 12-, and 27-fold increases, respectively, in the frequency of [risk](#)-related variants in the genes associated with Gaucher (GBA), Canavan (ASPA), and Tay Sachs (HEXA) among members of the AJ population. All three genes are associated with the brain and with other neurodegenerative diseases.

"This study was a global collaborative effort that will help us understand and hopefully improve how we treat Crohn's—but also quantifies the underlying increased risk for diseases such as Gaucher, Canavan, and Tay Sachs," said co-senior author Mark Daly, institute member and co-director of the Program in Medical and Population Genetics at the Broad, chief of Analytical and Translational Genetics Unit at Massachusetts General Hospital, and associate professor of medicine at Harvard Medical School. "It particularly highlights the value of using exome sequencing data across large [population](#) samples to determine risk factors."

**More information:** Manuel A. Rivas et al. Insights into the genetic epidemiology of Crohn's and rare diseases in the Ashkenazi Jewish population, *PLOS Genetics* (2018). [DOI: 10.1371/journal.pgen.1007329](https://doi.org/10.1371/journal.pgen.1007329)

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