



JAK3-deficient severe combined immunodeficiency

JAK3-deficient severe combined immunodeficiency (SCID) is an inherited disorder of the immune system. Individuals with *JAK3*-deficient SCID lack the necessary immune cells to fight off certain bacteria, viruses, and fungi. They are prone to repeated and persistent infections that can be very serious or life-threatening. Often the organisms that cause infection in people with *JAK3*-deficient SCID are described as opportunistic because they ordinarily do not cause illness in healthy people. Affected infants typically develop chronic diarrhea, a fungal infection in the mouth called oral thrush, pneumonia, and skin rashes. Persistent illness also causes affected individuals to grow more slowly than other children. Without treatment, people with *JAK3*-deficient SCID usually live only into early childhood.

Frequency

JAK3-deficient SCID accounts for an estimated 7 to 14 percent of cases of SCID. The prevalence of SCID from all genetic causes combined is approximately 1 in 50,000, although it may be higher in certain regions.

Causes

JAK3-deficient SCID is caused by mutations in the *JAK3* gene. The protein produced from this gene helps regulate the growth and maturation of certain types of white blood cells (lymphocytes) called T cells and natural killer cells. In addition, the *JAK3* protein is important for the normal maturation of another type of lymphocyte called B cells. T cells, B cells, and natural killer cells attack bacteria, viruses, and fungi, and help regulate the entire immune system.

Mutations in the *JAK3* gene prevent the production of *JAK3* protein or lead to production of a nonfunctional protein. A loss of functional *JAK3* protein results in the absence of T cells and natural killer cells and a normal number of poorly functioning B cells. This shortage of functional lymphocytes causes people with *JAK3*-deficient SCID to be susceptible to infections.

Inheritance Pattern

This condition is inherited in an autosomal recessive pattern, which means both copies of the gene in each cell have mutations. The parents of an individual with an autosomal recessive condition each carry one copy of the mutated gene, but they typically do not show signs and symptoms of the condition.

Other Names for This Condition

- autosomal recessive T-B+NK- SCID
- autosomal recessive T cell-negative, B cell-positive, NK cell-negative severe combined immunodeficiency
- JAK3 SCID
- T-B+ severe combined immunodeficiency due to JAK3 deficiency
- T cell-negative, B cell-positive, NK cell-negative SCID

Diagnosis & Management

Genetic Testing Information

- What is genetic testing?
[/primer/testing/geneticTesting](#)
- Genetic Testing Registry: Severe combined immunodeficiency, autosomal recessive, T cell-negative, B cell-positive, NK cell-negative
<https://www.ncbi.nlm.nih.gov/gtr/conditions/C1833275/>

Research Studies from ClinicalTrials.gov

- ClinicalTrials.gov
<https://clinicaltrials.gov/ct2/results?cond=%22JAK3+deficient+severe+combined+immunodeficiency%22+OR+%22JAK3+SCID%22+OR+%22Severe+Combined+Immunodeficiency%22>

Additional Information & Resources

Health Information from MedlinePlus

- Encyclopedia: Immunodeficiency disorders
<https://medlineplus.gov/ency/article/000818.htm>
- Health Topic: Immune System and Disorders
<https://medlineplus.gov/immunesystemanddisorders.html>

Genetic and Rare Diseases Information Center

- Severe combined immunodeficiency
<https://rarediseases.info.nih.gov/diseases/7628/severe-combined-immunodeficiency>

Additional NIH Resources

- National Human Genome Research Institute: Learning About Severe Combined Immunodeficiency (SCID)
<https://www.genome.gov/Genetic-Disorders/Severe-Combined-Immunodeficiency>
- National Institute of Allergy and Infectious Diseases: Severe Combined Immunodeficiency
<https://www.niaid.nih.gov/diseases-conditions/severe-combined-immunodeficiency-scid>

Educational Resources

- Great Ormond Street Hospital: Severe Combined Immunodeficiency (SCID)
<https://www.gosh.nhs.uk/conditions-and-treatments/conditions-we-treat/severe-combined-immunodeficiency-scid>
- KidsHealth from Nemours: Severe Combined Immunodeficiency (SCID)
<https://kidshealth.org/en/parents/severe-immunodeficiency.html>
- MalaCards: jak3-deficient severe combined immunodeficiency
https://www.malacards.org/card/jak3_deficient_severe_combined_immunodeficiency
- Merck Manual Consumer Version: Severe Combined Immunodeficiency (SCID)
<https://www.merckmanuals.com/home/immune-disorders/immunodeficiency-disorders/severe-combined-immunodeficiency-scid>
- Orphanet: T-B+ severe combined immunodeficiency due to JAK3 deficiency
https://www.orpha.net/consor/cgi-bin/OC_Exp.php?Lng=EN&Expert=35078

Patient Support and Advocacy Resources

- Immune Deficiency Foundation
<https://primaryimmune.org/>
- International Patient Organization for Primary Immunodeficiencies
<https://ipopi.org/>
- Jeffrey Modell Foundation
<http://www.info4pi.org/>
- National Organization for Rare Disorders
<https://rarediseases.org/rare-diseases/severe-combined-immunodeficiency/>

Scientific Articles on PubMed

- PubMed
<https://www.ncbi.nlm.nih.gov/pubmed?term=%28%28JAK3%5BTIAB%5D%29+AND+%28severe+combined+immunodeficiency%5BTIAB%5D%29%29+OR+%28%28JAK3%5BTIAB%5D%29+AND+%28SCID%5BTIAB%5D%29%29+OR+%28JAK3+severe+combined+immunodeficiency%5BTIAB%5D%29+AND+english%5Bla%5D+AND+human%5Bmh%5D+AND+%22last+3600+days%22%5Bdp%5D>

Catalog of Genes and Diseases from OMIM

- SEVERE COMBINED IMMUNODEFICIENCY, AUTOSOMAL RECESSIVE, T CELL-NEGATIVE, B CELL-POSITIVE, NK CELL-NEGATIVE
<http://omim.org/entry/600802>

Medical Genetics Database from MedGen

- Severe combined immunodeficiency, autosomal recessive, T cell-negative, B cell-positive, NK cell-negative
<https://www.ncbi.nlm.nih.gov/medgen/331474>

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