Y chromosome infertility

Y chromosome infertility is a condition that affects the production of sperm, making it difficult or impossible for affected men to father children. An affected man's body may produce no sperm cells (azoospermia), a smaller than usual number of sperm cells (oligospermia), or sperm cells that are abnormally shaped or that do not move properly.

Some men with Y chromosome infertility who have mild to moderate oligospermia may eventually father a child naturally. Assisted reproductive technologies may help other affected men; most men with Y chromosome infertility have some sperm cells in the testes that can be extracted for this purpose. The most severely affected men do not have any mature sperm cells in the testes. This form of Y chromosome infertility is called Sertoli cell-only syndrome.

Men with Y chromosome infertility usually do not have any other signs or symptoms. Occasionally they may have unusually small testes or undescended testes (cryptorchidism).

Frequency

Y chromosome infertility occurs in approximately 1 in 2,000 to 1 in 3,000 males of all ethnic groups. This condition accounts for between 5 percent and 10 percent of cases of azoospermia or severe oligospermia.

Causes

As its name suggests, this form of infertility is caused by changes in the Y chromosome. People normally have 46 chromosomes in each cell. Two of the 46 chromosomes are sex chromosomes, called X and Y. Females have two X chromosomes (46,XX), and males have one X chromosome and one Y chromosome (46,XY). Because only males have the Y chromosome, the genes on this chromosome tend to be involved in male sex determination and development.

Y chromosome infertility is usually caused by deletions of genetic material in regions of the Y chromosome called azoospermia factor (AZF) A, B, or C. Genes in these regions are believed to provide instructions for making proteins involved in sperm cell development, although the specific functions of these proteins are not well understood.

Deletions in the AZF regions may affect several genes. The missing genetic material likely prevents production of a number of proteins needed for normal sperm cell development, resulting in Y chromosome infertility.

In rare cases, changes to a single gene called USP9Y, which is located in the AZFA region of the Y chromosome, can cause Y chromosome infertility. The USP9Y gene provides instructions for making a protein called ubiquitin-specific protease 9. A small
number of individuals with Y chromosome infertility have deletions of all or part of the
USP9Y gene, while other genes in the AZF regions are unaffected. Deletions in the
USP9Y gene prevent the production of ubiquitin-specific protease 9 or result in the
production of an abnormally short, nonfunctional protein. The absence of functional
ubiquitin-specific protease 9 impairs the production of sperm cells, resulting in Y
chromosome infertility.

Inheritance Pattern

Because Y chromosome infertility impedes the ability to father children, this condition
is usually caused by new deletions on the Y chromosome and occurs in men with no
history of the disorder in their family. When men with Y chromosome infertility do father
children, either naturally or with the aid of assisted reproductive technologies, they
pass on the genetic changes on the Y chromosome to all their sons. As a result, the
sons will also have Y chromosome infertility. This form of inheritance is called Y-linked.
Daughters, who do not inherit the Y chromosome, are not affected.

Other Names for This Condition

• spermatogenic failure, Y-linked
• Y chromosome-related azoospermia

Diagnosis & Management

Genetic Testing Information

• What is genetic testing?
  /primer/testing/genetictesting
• Genetic Testing Registry: Spermatogenic failure, Y-linked 2
• Genetic Testing Registry: Spermatogenic failure, Y-linked, 1

Research Studies from ClinicalTrials.gov

• ClinicalTrials.gov
  https://clinicaltrials.gov/ct2/results?cond=%22Infertility%2C+Male%22+OR+%22Y
  +chromosome+infertility%22

Other Diagnosis and Management Resources

• GeneReview: Y Chromosome Infertility
  https://www.ncbi.nlm.nih.gov/books/NBK1339
• MedlinePlus Encyclopedia: Semen Analysis
  https://medlineplus.gov/ency/article/003627.htm
Additional Information & Resources

Health Information from MedlinePlus
- Encyclopedia: Semen Analysis
  https://medlineplus.gov/ency/article/003627.htm
- Health Topic: Assisted Reproductive Technology
  https://medlineplus.gov/assistedreproductivetechology.html
- Health Topic: Male Infertility
  https://medlineplus.gov/maleinfertility.html

Genetic and Rare Diseases Information Center
- Y chromosome infertility
  https://rarediseases.info.nih.gov/diseases/185/y-chromosome-infertility

Educational Resources
- American Society for Reproductive Medicine: Male Infertility
  https://www.reproductivefacts.org/topics/topics-index/male-infertility/
- MalaCards: y chromosome infertility
  https://www.malacards.org/card/y_chromosome_infertility

Patient Support and Advocacy Resources
- RESOLVE: The National Infertility Association
  https://resolve.org/infertility-101/medical-conditions/male-factor/

Clinical Information from GeneReviews
- Y Chromosome Infertility
  https://www.ncbi.nlm.nih.gov/books/NBK1339

Scientific Articles on PubMed
- PubMed
  https://www.ncbi.nlm.nih.gov/pubmed?term=%28y+chromosome+infertility%29+AND+english%5Bla%5D+AND+human%5Bmh%5D+AND+%22last+360+days+22%5Bdp%5D

Catalog of Genes and Diseases from OMIM
- SPERMATOCYTIC FAILURE, Y-LINKED, 1
  http://omim.org/entry/400042
- SPERMATOCYTIC FAILURE, Y-LINKED, 2
  http://omim.org/entry/415000
Sources for This Summary

  Citation on PubMed: https://www.ncbi.nlm.nih.gov/pubmed/18440997

  Citation on PubMed: https://www.ncbi.nlm.nih.gov/pubmed/20301513

Reprinted from Genetics Home Reference: 

Reviewed: January 2009
Published: October 9, 2018

Lister Hill National Center for Biomedical Communications
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