

# FREM2 Gene

Subjects: **Genetics & Heredity**

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FRAS1 related extracellular matrix protein 2

genes

## 1. Normal Function

The *FREM2* gene provides instructions for making a protein that is part of a group of proteins called the FRAS/FREM complex; in addition to being part of the complex, *FREM2* regulates the complex's formation. The FRAS/FREM complex is found in basement membranes, which are thin, sheet-like structures that separate and support cells in many tissues. The complex is particularly important during development before birth. One of its roles is to anchor the top layer of skin by connecting the basement membrane of the top layer to the layer of skin below. The FRAS/FREM complex is also involved in the proper development of certain other organs and tissues, including the kidneys, although the mechanism is unclear.

## 2. Health Conditions Related to Genetic Changes

### 2.1 Fraser Syndrome

At least two mutations in the *FREM2* gene have been found to cause Fraser syndrome; these mutations are involved in a small percentage of cases of this condition. Fraser syndrome affects development before birth and is characterized by eyes that are completely covered by skin (cryptophthalmos), fusion of the skin between the fingers and toes (cutaneous syndactyly), and abnormalities of the kidneys and other organs and tissues.

*FREM2* gene mutations involved in Fraser syndrome lead to production of an abnormal *FREM2* protein that likely does not function properly. As a result, the FRAS/FREM complex cannot form. Lack of the FRAS/FREM complex in the basement membrane of skin leads to detachment of the top layer of skin, causing blisters to form during development. These blisters likely prevent the proper formation of certain structures before birth, leading to cryptophthalmos and cutaneous syndactyly. It is unknown how lack of the FRAS/FREM complex leads to kidney abnormalities and other problems in Fraser syndrome.

### 2.2 Coloboma

### 2.3 Congenital Anomalies of Kidney and Urinary Tract

## 2.4 Other Disorders

Mutations in the *FREM2* gene have also been found in people with abnormalities of the kidneys and urinary tract but no other signs and symptoms of Fraser syndrome (described above). Such abnormalities are grouped together as congenital anomalies of the kidney and urinary tract (CAKUT). The *FREM2* gene mutations involved in CAKUT typically change single protein building blocks (amino acids) in the *FREM2* protein. Researchers speculate that the effects of these mutations are milder than those of mutations that cause Fraser syndrome; some *FREM2* protein function may still remain. How these gene mutations affect the FRAS/*FREM* complex or lead to renal agenesis and other CAKUT is unknown.

## 3. Other Names for This Gene

- DKFZp686J0811
- ECM3 homolog
- FRAS1-related extracellular matrix protein 2

## References

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