

Consent form - First in Class

Genomic testing is being offered to you/your child/your ward to determine the cause of your/their health problem. The information on this form briefly summarizes the information about genomic testing that was discussed with you by your healthcare provider. If you have any questions or concerns about this testing, please contact your healthcare provider to discuss further.

Purpose

Alamy Health, in collaboration with its partners, offers genomic testing (the “First in Class” test). This genomic testing involves whole genome sequencing that looks for changes or variations throughout one’s DNA. Such DNA variations can cause a gene or a group of genes to function improperly. Additionally, if indicated, testing might include assessment for DNA methylation alterations and alterations of the messages, aka RNA, encoded by genes. These analyses can provide valuable information about the cause of the health concerns and possible management options.

How it is performed

Genomic testing is performed on DNA that is usually obtained from a small blood sample. Analysis and reporting of the testing data are guided by detailed clinical information and family history information along with up to date published medical and scientific information. In some cases, samples from other family members are needed to interpret the results of the individual with health concerns (e.g., samples from parents along with a child’s sample). Alamy Health in partnership with Breakthrough Genomics will provide a test report to the healthcare provider who ordered the test. This healthcare provider will be responsible for discussing the result with the individual/family and guiding management. If relevant, the test report will indicate if a genetic change was inherited from one or both of the parents. However, reports are typically not issued for family members whose samples were used to guide interpretation for the individual with the health concerns, unless testing for secondary findings has been requested (see below section on “Secondary Findings”).

Types of results

Positive genetic result: A genetic change was found that is the likely cause of the health concerns. These genetic changes are termed pathogenic variants or likely pathogenic variants.

Positive epigenetic results: Two types of DNA methylation changes can be found in support of the pathogenicity (cause of the health concern) of certain genetic variants. One type is a methylation alteration that is localized to a specific genomic region.

The second type is a specific genome-wide pattern of DNA methylation alterations which constitutes a signature characteristic of a specific disorder. It is possible that a DNA methylation change indicative of a genetic disorder is detected without an associated genomic variant; in such instances, further testing might be indicated.

Positive transcriptome result: Sequencing of RNA provides support for the pathogenicity of the genetic variant.

Negative: No genetic, epigenetic, or transcriptomic changes were found to explain the health concerns. This might be due to current limitations in scientific and medical knowledge and/or testing technology. Such results do not, however, rule out the possibility of an underlying genetic disease. Reanalysis or retesting in future might be indicated and you should discuss this with your healthcare provider (see below section on “Variant(s) of uncertain significance” and “Sample retention, data storage, and database participation”).

Variant(s) of uncertain significance (VUS): Change(s) was/were found, but it is unclear whether this change is the cause of the health concerns. If relevant, changes in the epigenome and transcriptome might also be reported. The interpretation could remain uncertain until additional information becomes available. Such information could include published scientific articles indicating that the finding is pathogenic or that the finding does not cause health concerns (benign).

Secondary findings: The test might reveal findings, termed secondary findings. Such findings are not related to the current reasons for testing but could have important implications for current and/or future health. The American College of Medical Genetics (ACMG) has recommended the reporting of genetic variants in select genes associated with various genetic disorders that are considered to be medically actionable in childhood or in adulthood

(<https://www.acmg.net/PDFLibrary/Secondary-Findings-v3.1.pdf>). You can choose to have the ACMG list of secondary findings analyzed in yourself/your child/ward, and any pathogenic or likely pathogenic variants in these genes will be reported. Alternatively, you can decline to have the ACMG list of secondary findings analyzed in yourself/your child/ward. The decisions you make regarding the reporting of secondary findings in yourself/your child/ward will not impact the analysis and reporting of findings related to the presenting health concerns. The decision to report/not report secondary findings should be discussed with your healthcare provider.

Risks and limitations

- There may be standard risks associated with blood draw (e.g., bruising, bleeding).
- Inaccurate results can occur due to:
 - i) mislabeled samples regardless of strict clinical and laboratory protocols in place to minimize this risk
 - ii) inaccurate or incomplete reporting of medical and family history information
 - iii) rare technical errors
 - iv) other reasons
- This test might reveal biological relationships that were previously unrecognized, such as non-paternity or non-maternity.
- Genomic data may provide ancestry information (e.g. ethnicity) which could be used for the sole purpose of aiding the analysis of this test.
- Due to limitations in current medical knowledge or testing technology, a genetic variant may not be identified even though one exists and may contribute to or cause health problems.
- Genetic testing may identify the diagnosis for the health problems in you/your child/ward; however, the test cannot predict the severity and precise clinical course of the genetic disorder. Your healthcare provider can discuss these issues with you.
- Genetic test results can trigger difficult emotions and, in some cases, cause tension within a family since identifying a genetic diagnosis can have implications for other family members.
- Test results might impact insurance coverage and rates, as well as employment. There may be local laws that help protect individuals undergoing genetic testing against discrimination. Please consider discussing this with your healthcare provider.

Sample retention, data storage and database participation

After testing, Alamy Health will store the individual's biological sample, if available, and genomic data for the option of reanalysis and reinterpretation of the data in the future. Upon request by a health provider, Alamy Health will determine whether reanalysis or retesting would be preferred.

Alamy Health and its partners adhere to rigorous standards for personal health data storage as guided by laboratory regulatory bodies.

Alamy Health in collaboration with Breakthrough Genomics, will share de-identified genomic and phenotypic (i.e. clinical) information with select genomic data repositories in order to improve the medical and scientific community's understanding of genetic information. Although sharing such information could support my/my child's/my ward's diagnosis and health management, this could also result in no personal benefit. See notices of privacy practice for more details relative to the sharing of genetic information.

After testing is complete, an individual's sample might be de-identified and used for internal validation, quality improvement and product development.

Insurance and billing

Alamy Health's partner laboratory, Breakthrough Genomics, will bill either the patient's health insurance plan or the institution (i.e. hospital) as per the selection indicated on the test requisition. Alamy Health in collaboration with Breakthrough Genomics will be authorized to release any information required for billing. The patient is financially responsible for any costs that are not covered by their insurance.

Please check ONE of the following:

☐ YES, I DO wish for secondary findings to be reported in myself/my child/ward

☐ NO, I DO NOT wish for secondary findings to be reported in myself/my child/ward

<i>I have reviewed this consent form with my healthcare provider and have been given the opportunity to ask questions. By signing below, I consent to testing by Alamy Health's partner lab, Breakthrough Genomics, as described in this consent form.</i>	
Name (print)	Date of birth (MM-DD-YYYY)
Signature (if applicable)	Date (MM-DD-YYYY)
Name and relationship of legal decision maker to patient (print)	Signature of legal decision maker