CancerNext

GENETIC TESTING FOR HEREDITARY CANCER





What Are the Causes of Hereditary Cancer?

Familial Cancer

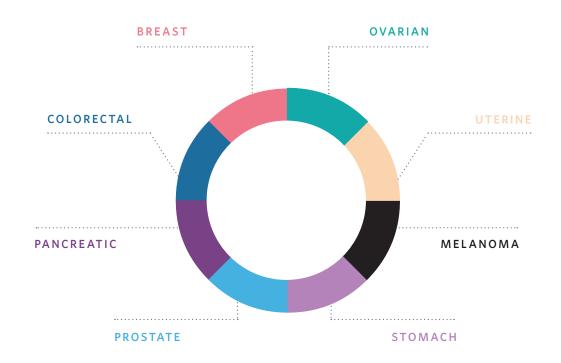
A clustering of cancer in a family that may be due to genes and/or other shared factors, such as environment and lifestyle

Hereditary Cancer

A clustering of cancer in a family due to inherited gene changes (mutations), which can be passed from parent to child



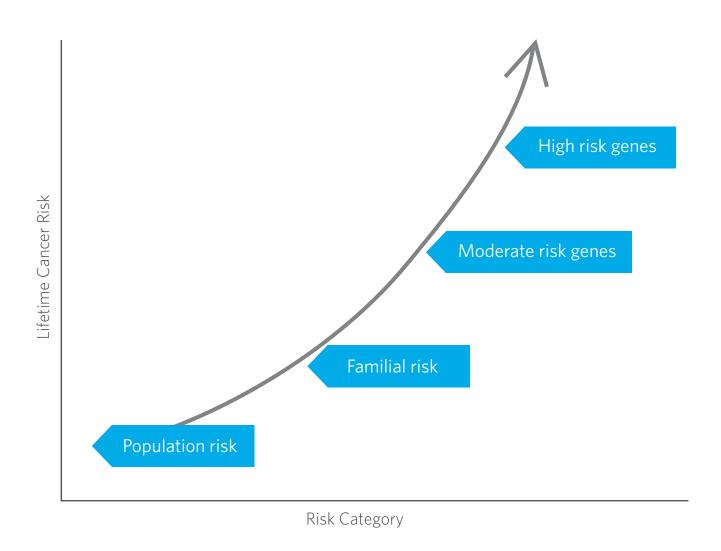
What Types of Cancer Can Be Hereditary*?





^{*}These are the most common types of cancer that can be hereditary and is not a comprehensive list.

What Are the Cancer Risks by Gene Type?





What Do the Cancer Gene Risk Levels Mean?

High Risk Genes

• Have published recommendations for cancer screening and/or prevention

Moderate Risk Genes

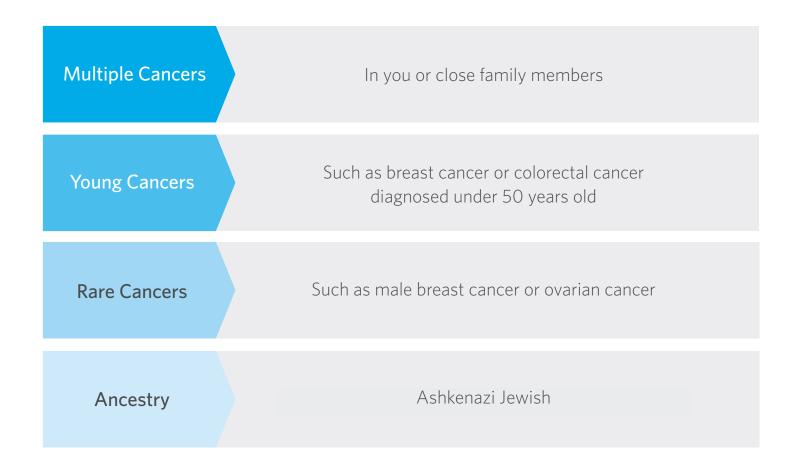
- May have newer recommendations for screening and/or prevention
- If no recommendations are available, screening may be based on family history and/or other risk factors

Newly Described Genes

- Are associated with an increased lifetime risk of developing certain types of cancer, but the risk is not well-defined
- Recommendations may be based on family history and/or other risk factors



What Are Signs of Hereditary Cancer?





CancerNext: High Risk Genes and Associated Cancers

GENES	BREAST	OVARIAN	COLORECTAL	UTERINE	PANCREATIC	PROSTATE	STOMACH	MELANOMA	OTHERS
APC			✓		✓				✓
BMPR1A			✓				√		
BRCA1	√	✓			\checkmark	✓			\checkmark
BRCA2	√	✓			\checkmark	✓		\checkmark	\checkmark
CDH1	\checkmark						\checkmark		\checkmark
CDKN2A					✓			✓	✓
CDK4								✓	✓
EPCAM		✓	\checkmark	✓	✓	✓	\checkmark		\checkmark
MLH1		\checkmark	\checkmark	✓	✓	\checkmark	\checkmark		\checkmark
MSH2		✓	✓	√	✓	✓	√		\checkmark
MSH6		✓	\checkmark	✓	✓	✓	\checkmark		\checkmark
MUTYH biallelic	\checkmark		\checkmark	✓					
PMS2		✓	\checkmark	✓	✓	✓	\checkmark		\checkmark
PTEN	\checkmark		✓	✓				\checkmark	\checkmark
SMAD4			✓				√		
STK11	√	✓	✓		✓				\checkmark
TP53	\checkmark	\checkmark	✓	√	✓	✓	\checkmark	\checkmark	\checkmark



CancerNext: Moderate Risk Genes and Associated Cancers

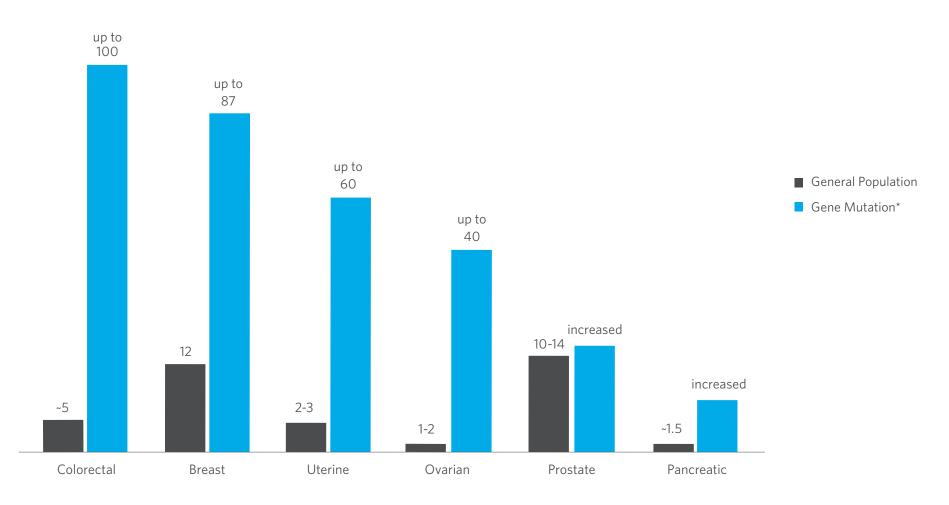
GENES	BREAST	OVARIAN	COLORECTAL	UTERINE	PANCREATIC	PROSTATE	STOMACH	MELANOMA	OTHERS
ATM	✓				✓	✓			
BRIP1	√	\checkmark							
CHEK2	√	\checkmark	\checkmark			\checkmark			\checkmark
GREM1			\checkmark						
NF1	√								\checkmark
PALB2	√	\checkmark			\checkmark	\checkmark			
POLD1			\checkmark						
POLE			✓						
RAD51C	✓	\checkmark							
RAD51D	√	✓				√			
TP53	✓	\checkmark	✓		✓	✓	√	✓	\checkmark



CancerNext: Newly Described Genes and Associated Cancers

GENES	BREAST	OVARIAN	COLORECTAL	UTERINE	PANCREATIC	PROSTATE	STOMACH	MELANOMA	OTHERS
BARD1	√								
DICER1		✓							\checkmark
HOXB13						✓			
MRE11A	\checkmark								
MUTYH Carrier	\checkmark		\checkmark						
NBN	\checkmark					✓			\checkmark
RAD50	\checkmark								
SMARCA4		\checkmark							\checkmark

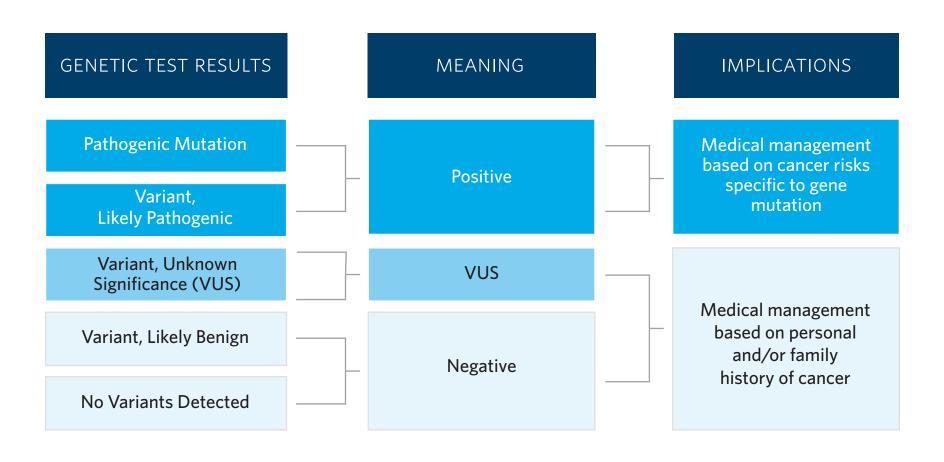
Lifetime Cancer Risks with CancerNext Gene Mutations (%)



^{*}Mutations in most CancerNext genes lead to an increased risk for some cancers in this graph, but not all. This graph represents the highest risks associated with some CancerNext genes.



What Do My Test Results Mean?





Positive Test Results

	HIGH RISK GENE	MODERATE RISK OR NEWLY DESCRIBED GENE				
Result	A mutation was found	A mutation was found				
Cancer Risks	High risk (for certain types of cancer)	Moderate risk: Moderately increased risk (for certain types of cancer) Newly described: Increased risk (for certain types of cancer), but exact risk is not yet defined				
Cancer Screening and Prevention	Recommendations based on genetic test result	Recommendations based on genetic test results and/or family history				
Genetic Testing For Family Members	Recommend genetic counseling and consideration of genetic testing	Recommend genetic counseling and consideration of genetic testing				



Negative Test Results

Result No mutations were found **Cancer Risks** Based on personal and/or family history **Cancer Screening** Recommendations based on personal and/or family and Prevention history **Genetic Testing For** May be considered in certain situations Family Members



Variant of Unknown Significance Test Results

A genetic change was found but it is unclear whether Result or not it causes increased risk for cancer Cancer Risks Based on personal and/or family history **Cancer Screening** Recommendations based on personal and/or family and Prevention history **Genetic Testing For** May be considered in certain situations **Family Members**



