

CENTOGENE  
THE RARE DISEASE COMPANY

آزمایشگاه دنا  
DeNA Laboratory

**CentoCancer<sup>®</sup>**

STRIVE FOR THE MOST  
COMPLETE INFORMATION

## **CentoCancer® – our most complete Oncogenetics panel for hereditary mutations**

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Certain hereditary pathogenic variants confer an increased risk of developing cancers during an individual's lifetime. The early identification of pathogenic variants in cancer predisposition genes represents a fundamental step in the diagnosis, management and treatment of individuals and families with hereditary cancer syndromes.

### **PANEL COMPOSITION**

CentoCancer®, our most comprehensive cancer panel has now been extended to 56 genes, offering you complete answers to help you choose the best possible therapeutic approach for your patients. Each gene in CentoCancer® has been carefully selected based on its risk potential in the development of one or more of the following cancers:

- Breast
- Ovarian
- Colorectal
- Gastric
- Bowel
- Endometrial
- Pancreatic
- Melanoma
- Renal
- Prostate

### **WHO SHOULD CONSIDER CentoCancer® FOR GENETIC TESTING?**

CentoCancer® is appropriate for:

- Individuals with a positive personal history of early-onset cancer, rare cancer, bilateral cancer, or multiple primary cancers
- Unaffected individuals with a positive family history of multiple generations of cancers, rare cancers or early-onset cancers
- Individuals in whom the suspected genetic diagnoses are not covered by a single targeted panel or if a targeted panel testing was previously negative

## Centocancer® – Panel composition and methodology

Centocancer® includes the following 56 most relevant cancer associated genes:

APC	CDH1	HNF1B	MSH2	POLD1	RAD51D	STK11
ATM	CDK4	HOXB13	MSH6	POLE	RET	TP53
BARD1	CDKN2A	MC1R	MUTYH	POT1	SDHA	TSC1
BLM	CHEK2	MEN1	NBN	PRSS1	SDHAF2	TSC2
BMPR1A	EPCAM	MET	NTHL1	PTCH1	SDHB	VHL
BRCA1	FH	MITF	PALB2	PTEN	SDHC	WT1
BRCA2	FLCN	MLH1	PMS1	RAD50	SDHD	XRCC2
BRIP1	HNF1A	MRE11A	PMS2	RAD51C	SMAD4	XRCC3

### KEY PANEL FACTS

- NGS bidirectional sequencing of all 56 genes in the panel, including coding regions (all exons) and exon/intron boundaries +/-10bp
- Coverage: >99% of target bases covered at >20x; mean coverage ≥180x
- 100% coverage of core genes: BRCA1, BRCA2, TP53
- CNV analysis from NGS data included for all genes
- All reported variants confirmed by Sanger sequencing or MLPA/qPCR
- All relevant deep intronic mutations described in HGMD® 2017.3 and CentoMD® included
- Turnaround Time: 10 business days
- Required Material: ≥4µg DNA or ≥1ml EDTA blood or ≥1 filter card

## Some common cancer predisposition syndromes covered by CentoCancer®

### Syndromes

#### HEREDITARY BREAST/OVARIAN CANCER

BRCA1, BRCA2

#### LI-FRAUMENI SYNDROME

TP53

#### COWDEN SYNDROME

PTEN

#### HNPCC (LYNCH SYNDROME)

MLH1, MSH2, MSH6, PMS1, PMS2

#### FAMILIAL ADENOMATOUS POLYPOSIS

APC

#### VON HIPPEL-LINDAU

VHL

#### MULTIPLE ENDOCRINE NEOPLASIA

MEN1, RET

#### RETINOBLASTOMA

RB1

### Associated cancers



Breast, ovarian, prostate, pancreatic, melanoma



Breast, sarcomas, adrenocortical carcinoma, leukemia, brain tumors



Breast, thyroid, benign lesions of skin, hamartoma, renal cell carcinoma, uterine



Colorectal (often right sided and multifocal), endometrial, ovarian, small bowel, stomach, pancreas, ureter, renal pelvis



Polyposis, colorectal, thyroid, gastric, periampullary carcinoma, hepatoblastoma



Renal cell carcinoma, retinal angioma, cerebellar hemangioblastoma, pheochromocytoma, pancreatic cysts, islet cell tumor



Parathyroid tumors, pancreatic tumors, pituitary tumors, medullary thyroid cancer, pheochromocytoma, neuromas



Retinoblastoma, often bilateral and <1 year of age, also associated increased risk of soft tissue sarcomas, melanoma, brain tumors

# HEREDITARY CANCER AND/OR SUSCEPTIBILITY

Selection of genetic test/panel according to family history and clinical data

BRCA1, BRCA2 panel	BRCA1, BRCA2
Breast ovarian cancer panel	CDH1, PTEN, STK11, TP53
Breast ovarian cancer panel PLUS	ATM, BARD1, BRIP1, CHEK2, MRE11, MLH1, MRE11A, MSH2, MSH6, MUTHY, NBN, PALB2, PMS1, PMS2, RAD50, RAD51C, RAD51D, XRCC2
Centobreast® panel	ATM, BARD1, BRCA1, BRCA2, BRIP1, CDH1, CHEK2, NBN, PALB2, PTEN, RAD51C, STK11, TP53
Colon cancer with polyps panel	APC, BMP11A, MUTHY, PTEN, SMAD4, STK11
Colon cancer non-polyposis panel	EPCAM, MSH2, MLH1, MSH6, PMS2
Centocolon extended panel	APC, BMP11A, CDH1, CHEK2, EPCAM, MLH1, MSH2, MSH6, MUTHY, NTHL1, PMS2, FOLD1, POLE, PTEN, SMAD4, STK11, TP53
Gastric cancer panel, targeted	BMP11A, CDH1, EPCAM, MLH1, MSH2, MSH6, PMS1, PMS2, SMAD4
Ovarian cancer panel, targeted	BARD1, BRCA1, BRCA2, BRIP1, EPCAM, MLH1, MRE11A, MSH2, MSH6, NBN, PMS1, PMS2, RAD50, RAD51C, RAD51D, STK11, TP53
Prostate cancer panel	BRCA1, BRCA2, CHEK2, HMOX1, MLH1, MSH2, MSH6, NBN, PTEN, TP53
Pancreatic cancer panel, targeted	APC, ATM, BMP11A, CDKN2A, EPCAM, MLH1, MSH2, MSH6, PALB2, PMS1, PMS2, PRSS1, SMAD4, STK11
Renal cancer panel, targeted	EPCAM, FH, FLCN, HNF1A, HNF1B, MET, MTF, MLH1, MSH2, MSH6, PMS1, PMS2, PTEN, SDHB
Skin cancer panel, targeted	CDKN2A, EPCAM, MC1R, MTR, MLH1, MSH2, MSH6, PMS1, PMS2, POT1, PTCN1, XRCC3
Thyroid cancer panel, targeted	APC, PTEN, RET
Uterine cancer panel, targeted	EPCAM, MLH1, MSH2, MSH6, PMS1, PMS2, PTEN
PGL / PCC / GIST panel, targeted	GNAS, NF1B, MAX, MEN1, NF1, RET, SDHA, SDHAF2, SDHB, SDHC, SDHD, TMEM127, TP53, VHL
Multiple endocrine neoplasias / paraganglioma/pheochromocytoma panel	CDKN1B, MAX, MEN1, RET, SDHA, SDHAF2, SDHB, SDHC, SDHD, TMEM127, VHL

Identification of specific cancer-causing pathogenic variant

Genetic counseling, genetic testing of all family members with consent

No pathogenic variants identified

Complex family history, variability of cancers and absence of known genetic cause in the family

Centocancer® panel	APC, ATM, BARD1, BLN1, BMP11A, BRCA1, BRCA2, BRIP1, CDH1, CDKN2A, CDKN2A, CHEK2, EPCAM, FH, FLCN, HNF1A, HNF1B, HMOX1, MC1R, MEN1, MET, MTR, MLH1, MRE11A, MSH2, MSH6, MUTHY, NBN, NTHL1, PALB2, PMS1, PMS2, FOLD1, POLE, POT1, PRSS1, PTCN1, PTEN, RAD50, RAD51C, RAD51D, RET, SDHA, SDHAF2, SDHB, SDHC, SDHD, SMAD4, STK11, TP53, TSC1, TSC2, VHL, WT1, XRCC2, XRCC3
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Identification of specific cancer-causing pathogenic variant

Genetic counseling, genetic testing of all family members with consent

No pathogenic variant identified

WES analysis

Identification of specific cancer-causing pathogenic variants

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