

# **Supplemental Material**

## **BRCA2 Polymorphisms and Breast Cancer Susceptibility: a Multi-Tools Bioinformatics Approach**

Haris Jan<sup>a</sup> Najeeb Ullah Khan<sup>a</sup> Ayman M. Al-Qaaneh<sup>b</sup> Munazzah Tasleem<sup>c</sup> Mikhlid H. Almutairi<sup>d</sup>  
Ijaz Ali<sup>e</sup>

<sup>a</sup>Institute of Biotechnology & Genetic Engineering (Health Division), The University of Agriculture Peshawar, Pakistan, <sup>b</sup>Department of Allied Health Sciences, Faculty of Nursing, Al-Balqa Applied University (BAU), Al-Salt 19117, Jordan, <sup>c</sup>Department of Biochemistry, Jamia Hamdard, Delhi, India, <sup>d</sup>Zoology Department, College of Science, King Saud University, P.O. Box: 2455, 11451, Riyadh, Saudi Arabia,  
<sup>e</sup>Centre for Applied Mathematics and Bioinformatics (CAMB), Gulf University for Science and Technology, Hawally, Kuwait.

**Table 1.** List of deleterious missense SNPs in the BRCA2 gene using consensus bioinformatics tool.

rs-id	Variant ID	AA Position	SIFT	PolyPhen-2	PHD-SNP	Predict SNP	SNAP	ClicVar/Varsome
rs80358597	32890631-T-G	F12V	D	D	D	D	D	VUS
rs1370260227	32890653-G-T	C19F	D	D	D	D	D	VUS
rs774521832	32893214-A-G	D23G	D	D	D	D	D	VUS
rs80358961	32893219-G-A	G25R	D	D	D	D	D	VUS
rs397508045	32893238-G-T	W31L	D	D	D	D	D	VUS
rs1566215918	32893367-T-C	L74P	D	D	D	D	D	VUS
rs28897701	32893369-G-C	A75P	D	D	D	D	D	Benign
rs774152844	32893427-C-A	P94H	D	D	D	D	D	Benign
rs80358660	32899312-T-G	L139R	D	D	D	D	D	VUS
rs587782795	32900240-C-G	P143R	D	D	D	D	D	VUS
rs80358726	32900405-C-A	P168T	D	D	D	D	D	Benign
rs80358761	32900658-T-C	I180T	D	D	D	D	D	VUS
rs1334767632	32906574-T-C	L320P	D	D	D	D	D	VUS
rs80358412	32906798-T-G	W395G	D	D	D	D	D	VUS
rs398122730	32907348-G-A	G578D	D	D	D	D	D	VUS
rs397507275	32907371-A-G	K586E	D	D	D	D	D	VUS
rs772156559	32907390-A-G	Y592C	D	D	D	D	D	VUS
rs587780646	32907453-T-G	L613R	D	D	D	D	D	Benign
rs80358479	32907504-C-T	T630I	D	D	D	D	D	Benign
rs759389988	32910885-T-G	L798R	D	D	D	D	D	Benign
rs80358530	32911263-A-T	N924I	D	D	D	D	D	Benign
rs141702094	32911590-A-T	D1033V	D	D	D	D	D	VUS
rs558973276	32911842-T-C	I1117T	D	D	D	D	D	VUS
rs532871047	32911857-G-A	G1122E	D	D	D	D	D	VUS

**Table 2.** Prediction of the molecular mechanism by mutpred2

Mutation	MutPred2 score	Predicted molecular mechanism (P-value)
F12V	0.544	Altered Disordered interface (P = 0.03)
		Altered Transmembrane protein (P = 0.04)
		Altered Disordered interface (P = 0.03)
C19F	0.777	Loss of Relative solvent accessibility (P = 0.01)
		Altered Transmembrane protein (P = 0.04)
		Altered Disordered interface (P = 0.03)
D23G	0.672	Loss of Relative solvent accessibility (P = 0.02)
		Altered Disordered interface (P = 0.03)
		Gain of Intrinsic disorder (P = 0.05)
G25R	0.686	Gain of Helix (P = 0.03)
		Gain of Intrinsic disorder (P = 0.03)
		Altered Disordered interface (P = 0.03)
W31L	0.779	Altered Ordered interface (P = 0.01)
		Gain of Intrinsic disorder (P = 0.03)
		Altered Disordered interface (P = 0.03)
L74P	0.511	Altered Disordered interface (P = 0.03)
		Loss of Helix (P = 0.03)
P168T	0.581	-
		Gain of Intrinsic disorder (P = 0.03)
I180T	0.519	Altered Disordered interface (P = 0.03)
		Altered Metal binding (P = 0.01)
		Altered Ordered interface (P = 0.01)
		Altered Disordered interface (P = 0.03)
		Altered Metal binding (P = 0.01)
Y592C	0.609	Altered Ordered interface (P = 0.02)
		Altered DNA binding (P = 0.01)
		Loss of Methylation at K589 (P = 0.01)
		Altered Stability (P = 0.04)
D1033V	0.641	Loss of Acetylation at K1032 (P = 0.03)
F1219V	0.694	-
G1529R	0.693	Loss of Acetylation at K1530 (P = 0.03)
		Altered Disordered interface (P = 0.02)
		Altered DNA binding (P = 0.02)
A2231P	0.739	Gain of Intrinsic disorder (P = 0.04)
		Loss of Ubiquitylation at K2232 (P = 0.05)
		Loss of Methylation at K2232 (P = 0.03)
H2324P	0.647	Altered Disordered interface (P = 0.05)
		Altered Metal binding (P = 0.05)
C2332R	0.702	Gain of Helix (P = 0.02)
		Gain of Strand (P = 0.04)
		Loss of Loop (P = 0.03)
G2596R	0.769	Loss of B-factor (P = 0.04)
		Altered Transmembrane protein (P = 0.01)
C2605Y	0.506	Altered Transmembrane protein (P = 0.02)

