

Table S3. Description of Candidate Germline Variants from Multiplexed, Panel Sequencing of FIGC

Position GRCh38			Frequency Cohorts (nr patients)										Frequency 1000Genomes							
Chr	Coordinate	Patient Genotype	Gene	Transcript ID	cDNA	Type of Variant/Location	Protein	Impact	FIGC (n=50)	HDGC (n=17)	Sporadic GC (n=47)	Variation ID	TSI (n=107)	FIN (n=99)	GBR (n=91)	IBS (n=107)	Global AF	EUR AF	EAS AF	gnomAD
13	32337410	C/G	BRCA2	NM_000059.3	c.3055C>G	Missense	p.Leu1019Val	Moderate	1	0	0	rs55638633	1	0	0	0	2,00E-04	1,00E-03	0,00E+00	2,00E-04
1	45331485	G/T	MUTYH	NM_001048171.1	c.1216C>A	Missense	p.Leu420Met	Moderate	1	0	0	rs144079536	0	0	0	0	2,00E-04	1,00E-03	0,00E+00	2,00E-04
11	108289671	C/T	ATM	NM_000051.3	c.4306C>T	Missense	p.His1436Tyr	Moderate	1	0	0	rs544891616	1	0	0	0	2,00E-04	1,00E-03	0,00E+00	3,26E-05
8	16155081	C/T	MSR1	NM_138715.2	c.881G>A	Missense	Gly294Glu	Moderate	1	0	0	rs41440349	0	0	0	0	3,99E-04	0,00E+00	0,00E+00	4,00E-04
8	16168606	G/T	MSR1	NM_138715.2	c.482C>A	Missense	p.Thr161Asn	Moderate	1	0	0	rs76147566	1	0	0	0	3,99E-04	1,00E-03	1,00E-03	4,00E-04
22	44726610	C/T	PRR5	NM_001017528.2	c.271C>T	Missense	p.Arg91Trp	Moderate	1	0	0	rs201344303	0	0	0	0	2,00E-04	0,00E+00	0,00E+00	2,00E-04
2	189795860	C/T	PMS1	NM_000534.4	c.224C>T	Missense	p.Thr75Ile	Moderate	1	0	0	rs61756360	0	0	0	0	3,99E-04	0,00E+00	0,00E+00	4,00E-04
10	87965825	C/T	PTEN	NM_000314.6	c.*353C>T	3'UTR		Modifier	1	0	0	rs181234898	0	0	0	0	4,99E-03	1,00E-03	7,90E-03	5,00E-03
10	87966260	T/C	PTEN	NM_000314.6	c.*788T>C	3'UTR		Modifier	1	0	0	rs138309082	0	0	0	0	4,99E-03	1,00E-03	7,90E-03	5,00E-03
10	87967174	G/A	PTEN	NM_000314.6	c.*1702G>A	3'UTR		Modifier	1	0	0	rs150265244	0	0	0	0	4,99E-03	1,00E-03	7,90E-03	5,00E-03
10	87967417	G/A	PTEN	NM_000314.6	c.*1945G>A	3'UTR		Modifier	1	0	0	rs567800059	0	0	0	0	4,99E-03	1,00E-03	7,90E-03	5,00E-03
10	87967645	C/T	PTEN	NM_000314.6	c.*2173C>T	3'UTR		Modifier	1	0	0	rs186996550	0	0	0	0	7,99E-04	0,00E+00	0,00E+00	8,00E-04
1	27355895	C/T	MAP3K6	NM_004672.4	c.3711+131G>A	3'UTR		Modifier	1	0	0	rs554613210	0	0	0	0	2,00E-04	0,00E+00	0,00E+00	2,00E-04
10	86924090	C/T	BMPR1A	NM_004329.2	c.*371C>T	3'UTR		Modifier	35	0	0	rs550209042	0	0	0	0	2,00E-04	0,00E+00	0,00E+00	2,00E-04
3	36993862	C/T	MLH1	NM_001167617	c.-202C>T	5'UTR		Modifier	1	0	0	rs561267247	0	0	0	0	2,00E-04	0,00E+00	1,00E-03	2,00E-04
22	44676975	G/A	PRR5	NM_001017528.2	c.-11+8170G>A	5'UTR		Modifier	1	0	0	rs552907174	0	0	0	0	2,00E-04	0,00E+00	0,00E+00	2,00E-04
5	112775612	T/A	APC	NM_000038.5	c.423-17T>A	Intronic		Modifier	1	0	0	rs534684461	0	0	0	0	3,19E-03	1,00E-03	6,00E-03	3,20E-03
5	147828151	T/A	SPINK1	NM_003122.4	c.88-23A>T	Intronic		Modifier	1	0	0	rs199929811	0	0	0	0	3,99E-04	2,00E-03	0,00E+00	4,00E-04
18	51048865	G/A	SMAD4	NM_005359.5	c.424+5G>A	Intronic		Low	1	0	0	rs200772603	0	0	0	0	2,00E-04	0,00E+00	0,00E+00	2,00E-04
10	86899915	G/A	BMPR1A	NM_004329.2	c.430+25G>A	Intronic		Modifier	1	0	0	rs200416589	1	0	0	0	2,80E-03	1,00E-03	0,00E+00	2,80E-03
10	87925641	G/A	PTEN	NM_000314.6	c.209+84G>A	Intronic		Modifier	1	0	0	rs185262832	0	0	0	0	4,99E-03	1,00E-03	7,90E-03	5,00E-03
1	27362613	C/T	MAP3K6	NM_004672.4	c.1255+28G>A	Intronic		Modifier	1	0	0	rs538360379	1	0	0	0	2,00E-04	1,00E-03	0,00E+00	2,00E-04
2	47791206	T/C	MSH6	NM_000179.2	c.457+83T>C	Intronic		Modifier	1	0	0	rs150046242	0	0	0	0	2,00E-04	0,00E+00	0,00E+00	2,00E-04
18	51049362	G/C	SMAD4	NM_005359.5	c.454+38G>C	Intronic		Modifier	1	0	0	rs201924714	0	0	0	0	2,00E-04	0,00E+00	0,00E+00	2,00E-04
16	86512849	C/T	FOXF1	NM_001451.2	c.980-76C>T	Intronic		Modifier	2	0	0	rs139488479	1	0	0	0	3,99E-04	1,00E-03	0,00E+00	4,00E-04
11	108280904	G/T	ATM	NM_000051.3	c.3403-91G>T	Intronic		Modifier	1	0	0	rs372950664	1	0	0	0	7,99E-04	1,00E-03	0,00E+00	8,00E-04
7	117666855	C/T	CFTR	NM_000492.3	c.4243-53C>T	Intronic		Modifier	1	0	0	rs185664216	1	0	0	0	4,59E-03	1,00E-03	0,00E+00	4,60E-03
7	142751675	G/C	PRSS1	NM_002769.4	c.201-99G>C	Intronic		Modifier	1	0	0	rs530207004	1	0	0	0	2,00E-04	1,00E-03	0,00E+00	2,00E-04
7	142750586	C/T	PRSS1	NM_002769.4	c.72C>T	Synonymous	p.Ile24=	Low	3	0	0	rs372637371	0	0	0	0	3,99E-04	0,00E+00	2,00E-03	4,00E-04
17	43097246	G/A	BRCA1	NM_007294.3	c.591C>T	Synonymous	p.Cys197=	Low	1	0	0	rs1799965	1	0	0	0	3,99E-04	2,00E-03	0,00E+00	4,00E-04
10	86919293	C/T	BMPR1A	NM_004329.2	c.990C>T	Synonymous	p.Ala330=	Low	1	0	0	rs199808362	1	0	0	0	2,00E-04	1,00E-03	0,00E+00	2,00E-04

VUS - Variants of Uncertain Significance; NA - Not Available; na - not applicable

ExAC	ClinVar clinical significance (variation ID)	Selected for Sanger Validation	Validated by Sanger sequencing	SIFT	PolyPhen	FATHMM	COSMIC	
							ID	Detected as Somatic in:
1.26E-04	Benign, Likely Benign (37817)	No	na	Tolerated	Benign	NA	NA	NA
5.73E-04	VUS (41752)	Yes	Yes	Deleterious	Benign	NA	NA	NA
5.06E-05	VUS (187606)	Yes	Yes	Deleterious	Benign	NA	NA	NA
7.01E-04	NA	Yes	Yes	Deleterious	Probably damaging	NA	NA	NA
3.54E-04	NA	Yes	Yes	Tolerated	Benign	NA	NA	NA
6.98E-05	NA	Yes	Yes	Deleterious	Probably damaging	NA	NA	NA
6.22E-04	NA	Yes	Yes	Deleterious	Probably damaging	Pathogenic (score 0.99)	COSM6198026	Gastric Cancer
NA	Likely Benign (301429)	Yes	Yes			NA	NA	NA
NA	Likely Benign (301440)	Yes	Yes			NA	NA	NA
NA	Likely Benign (301472)	Yes	Yes			NA	NA	NA
NA	Likely Benign (301475)	Yes	Yes			NA	NA	NA
NA	Likely Benign (301482)	Yes	Yes			NA	NA	NA
NA	NA	Yes	Yes			NA	NA	NA
NA	NA	Yes	No			NA	NA	NA
NA	NA	Yes	Yes			NA	NA	NA
NA	NA	Yes	Yes			NA	NA	NA
1.43E-03	Likely benign (217981)	No	na			NA	NA	NA
2.51E-03	Benign, Likely Benign (239508)	No	na			NA	NA	NA
2.08E-04	VUS (127950)	Yes	Yes			NA	NA	NA
1.73E-03	NA	No	na			NA	NA	NA
NA	NA	No	na			NA	NA	NA
1.32E-05	NA	No	na			NA	NA	NA
NA	NA	No	na			NA	NA	NA
2.05E-04	NA	No	na			NA	NA	NA
NA	NA	No	na			NA	NA	NA
NA	NA	No	na			NA	NA	NA
NA	NA	No	na			NA	NA	NA
NA	NA	Yes	Yes			NA	NA	NA
1.74E-03	NA	No	na			NA	NA	NA
1.54E-03	Benign, Likely Benign (55642)	No	na			Pathogenic (score 0.80)	COSM5020112	Haemangioblastoma
4.07E-06	Likely Benign (184817)	No	na			NA	COSM1559326, COSM35609	Glioma