

eTable 2. Seventy-six distinct cysteine-altering *NOTCH3* mutations present in the patient cohort.

Protein change	Genomic change	Codon change	Exon	EGFR domain	Frequency
p.Cys43Gly	g.15197570A>C	c.127T>G	2	1	1
p.Arg54Cys	g.15197537G>A	c.160C>T	2	1	4
p.Cys55Ser	g.15197533C>G	c.164G>C	2	1	1
p.Cys65Tyr	g.15197503C>T	c.194G>A	2	1	1
p.Cys67Ser	g.15192518A>T	c.199T>A	3	1	1
p.Cys76Arg	g.15192491A>G	c.226T>C	3	1	1
p.Cys82Phe	g.15192472C>A	c.245G>T	3	2	1
p.Gly85Cys	g.15192464C>A	c.253G>T	3	2	1
p.Arg90Cys	g.15192449G>A	c.268C>T	3	2	27
p.Cys93Tyr	g.15192439C>T	c.278G>A	3	2	3
p.Arg110Cys	g.15192389G>A	c.328C>T	3	2	14
p.Ser126Cys	g.15192263T>A	c.376A>T	4	3	1
p.Cys128Phe	g.15192256C>A	c.383G>T	4	3	2
p.Arg133Cys	g.15192242G>A	c.397C>T	4	3	13
p.Cys134Tyr	g.15192238C>T	c.401G>A	4	3	2
p.Arg141Cys	g.15192218G>A	c.421C>T	4	3	77
p.Phe142Cys	g.15192214A>C	c.425T>G	4	3	1
p.Cys144Tyr	g.15192208C>T	c.431G>A	4	3	2
p.Tyr150Cys	g.15192190T>C	c.449A>G	4	3	1
p.Arg153Cys	g.15192182G>A	c.457C>T	4	3	39
p.Ser154Cys	g.15192179T>A	c.460A>T	4	3	1
p.Arg169Cys	g.15192134G>A	c.505C>T	4	4	72
p.Arg182Cys	g.15192095G>A	c.544C>T	4	4	69
p.Cys183Arg	g.15192092A>G	c.547T>C	4	4	1
p.Cys185Ser	g.15192086A>T	c.553T>A	4	4	1
p.Tyr189Cys	g.15192073T>C	c.566A>G	4	4	2
p.Cys194Ser	g.15192059A>T	c.580T>A	4	4	9
p.Cys194Arg	g.15192059A>G	c.580T>C	4	4	1
p.Cys194Ser	g.15192058C>G	c.581G>C	4	4	2
p.Cys201Arg	g.15192038A>G	c.601T>C	4	5	1
p.Arg207Cys	g.15192020G>A	c.619C>T	4	5	11
p.Cys212Arg	g.15192005A>G	c.634T>C	4	5	2
p.Cys222Arg	g.15191975A>G	c.664T>C	4	5	1
p.Cys224Tyr	g.15191968C>T	c.671G>A	4	5	6
p.Cys233Arg	g.15191850A>G	c.697T>C	5	5	1
p.Cys240Tyr	g.15191828C>T	c.719G>A	5	6	1
p.Cys245Ser	g.15191814A>T	c.733T>A	5	6	4
p.Cys245Arg	g.15191814A>G	c.733T>C	5	6	1

p.Cys251Arg	g.15191796A>G	c.751T>C	5	6	11
p.Tyr258Cys	g.15191774T>C	c.773A>G	5	6	1
p.Cys323Ser	g.15191493A>T	c.967T>A	6	8	2
p.Cys323Tyr	g.15191492C>T	c.968G>A	6	8	1
p.Arg332Cys	g.15191466G>A	c.994C>T	6	8	18
p.Cys355Ser	g.15189401C>G	c.1064G>C	7	9	1
p.Cys379Phe	g.15189329C>A	c.1136G>T	7	9	2
p.Cys402Ser	g.15189162C>G	c.1205G>C	8	10	1
p.Cys408Gly	g.15189145A>C	c.1222T>G	8	10	1
p.Arg421Cys	g.15189106G>A	c.1261C>T	8	10	3
p.Arg427Cys	g.15189088G>A	c.1279C>T	8	10	2
p.Cys428Arg	g.15189085A>G	c.1282T>C	8	10	2
p.Cys440Gly	g.15189049A>C	c.1318T>G	8	11	3
p.Arg449Cys	g.15189022G>A	c.1345C>T	8	11	3
p.Cys457Ser	g.15188997C>G	c.1370G>C	8	11	3
p.Cys466Tyr	g.15188330C>T	c.1397G>A	9	11	1
p.Ser476Cys	g.15188301T>A	c.1426A>T	9	12	1
p.Cys493Arg	g.15188250A>G	c.1477T>C	9	12	1
p.Cys504Gly	g.15187977A>C	c.1510T>G	10	12	1
p.Cys516Phe	g.15187940C>A	c.1547G>T	10	13	2
p.Cys522Ser	g.15187922C>G	c.1565G>C	10	13	1
p.Arg558Cys	g.15187273G>A	c.1672C>T	11	14	7
p.Arg578Cys	g.15187213G>A	c.1732C>T	11	14	1
p.Arg592Cys	g.15187171G>A	c.1774C>T	11	15	1
p.Cys597Arg	g.15187156A>G	c.1789T>C	11	15	1
p.Cys597Ser	g.15187155C>G	c.1790G>C	11	15	1
p.Arg607Cys	g.15187126G>A	c.1819C>T	11	15	12
p.Arg640Cys	g.15186911G>A	c.1918C>T	12	16	1
p.Gly953Cys	g.15181098C>A	c.2857G>T	18	24	1
p.Arg985Cys	g.15181002G>A	c.2953C>T	18	25	4
p.Arg1006Cys	g.15180807G>A	c.3016C>T	19	26	1
p.Tyr1021Cys	g.15180761T>C	c.3062A>G	19	26	1
p.Cys1061Tyr	g.15180217C>T	c.3182G>A	20	27	1
p.Arg1100Cys	g.15180101G>A	c.3298C>T	20	28	1
p.Cys1119Tyr	g.15179468C>T	c.3356G>A	21	28	4
p.Cys1222Gly	g.15179079A>C	c.3664T>G	22	31	2
p.Arg1231Cys	g.15179052G>A	c.3691C>T	22	31	5
p.Cys1250Gly	g.15178912A>C	c.3748T>G	23	32	1