

Supplementary Table 1 APC mutational data from the COSMIC database

Amino acid change	Nucleotide mutation	Relative mutation distribution in carcinomas, <i>n</i> (%)	Absolute mutation frequency in carcinomas, %
p.Q1291*	c.3871C>T	7 (0.68)	0.25
p.T1301fs*15	c.3900_3901insT	0 (0)	0
p.L1302fs*3	c.3903delC	1 (0.10)	0.04
p.I1304fs*4	c.3912delA	1 (0.10)	0.04
p.E1306*	c.3916G>T	8 (0.78)	0.29
	c.3921_3925delAA		
p.E1309fs*4	AAG	44 (4.30)	1.59
p.E1309*	c.3925G>T	12 (1.17)	0.43
p.K1310*	c.3928A>T	2 (0.20)	0.07
p.S1315*	c.3944C>A	4 (0.39)	0.14
<i>p.E1317Q</i>	c.3949G>C	2 (0.20)	0.07
p.V1320fs*11	c.3957_3958insT	0 (0)	0
p.E1322*	c.3964G>T	7 (0.68)	0.25
p.R1331*	c.3991A>T	2 (0.20)	0.07
p.Q1338*	c.4012C>T	12 (1.17)	0.43
p.S1344*	c.4031C>A	0 (0)	0
p.E1353*	c.4057G>T	6 (0.59)	0.22
p.S1356*	c.4067C>G	3 (0.29)	0.11
p.Q1367*	c.4099C>T	14 (1.37)	0.51
p.Q1378fs*7	c.4131_4132insT	0 (0)	0
p.Q1378*	c.4132C>T	11 (1.07)	0.40
p.E1397fs*1	c.4184_4185insT	0 (0)	0
p.S1400*	c.4199C>A	3 (0.29)	0.11
	c.4216_4217insCGT		
p.Q1406fs*11	TC	0 (0)	0
p.Q1406*	c.4216C>T	6 (0.59)	0.22
p.E1408*	c.4222G>T	5 (0.49)	0.18

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	c.4232_4238delGTG		
p.S1411fs*1	GAAT	0 (0)	0
p.S1411fs*4	c.4233delT	6 (0.59)	0.22
p.V1414fs*5	c.4241delT	0 (0)	0
p.G1416fs*3	c.4247delG	1 (0.10)	0.04
	c.4259_4272delCCC		
p.P1420fs*2	AGTGATCTTCC	0 (0)	0
p.S1421fs*52	c.4263delT	2 (0.20)	0.07
p.Q1429*	c.4285C>T	8 (0.78)	0.29
p.T1430fs*43	c.4287delC	0 (0)	0
	c.4294_4314delCAC		
	C		
	AAGCAGAAGTA		
p.P1432fs*35	AAAC	0 (0)	0
p.T1438fs*35	c.4312delA	4 (0.39)	0.14
p.T1438fs*35	c.4313delC	0 (0)	0
p.P1439fs*34	c.4316delC	1 (0.10)	0.04
p.P1441fs*32	c.4322delC	0 (0)	0
<i>p.P1442P</i>	<i>c.4326T&gt;A</i>	2 (0.20)	0.07
p.T1445fs*28	c.4333delA	0 (0)	0
p.T1445fs*28	c.4334delC	0 (0)	0
p.R1450*	c.4348C>T	60 (5.86)	2.17
p.E1461*	c.4381G>T	1 (0.10)	0.04
p.E1461fs*7	c.4382_4383insA	0 (0)	0
p.S1465fs*3	c.4386_4387delGA	19 (1.86)	0.69
p.K1462fs*10	c.4386delG	0 (0)	0
<i>p.P1483S</i>	<i>c.4447C&gt;T</i>	0 (0)	0
<i>p.A1485T</i>	<i>c.4453G&gt;A</i>	0 (0)	0
	c.4460_4464delCTT		
p.T1487fs*27	TA	0 (0)	0
p.L1488fs*19	c.4464delA	4 (0.39)	0.14

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p.H1490fs*17	c.4469_4470insTA	0 (0)	0
p.A1492fs*22	c.4473_4474insT	2 (0.20)	0.07
p.E1494fs*13	c.4480delG	3 (0.29)	0.11
<i>p.G1499R</i>	<i>c.4495G&gt;A</i>	0 (0)	0
other mutations within MCR			
(1286-1513)		477 (46.59)	17.24
Total mutants (MCR)		740 (72.27)	26.75
Total mutants (COSMIC)		1024 (100)	37.01

**Supplementary Table 2 *BRAF* mutational data from the COSMIC database**

<b>Amino acid change</b>	<b>Nucleotide mutation</b>	<b>Relative mutation distribution in carcinomas, <i>n</i> (%)</b>	<b>Absolute mutation frequency in carcinomas, %</b>
p.D594G	c.1781A>G	11 (0.37)	0.04
p.V600E	c.1799T>A	2862 (97.38)	10.05
p.V600M	c.1798G>A	1 (0.03)	0
p.K601E	c.1801A>G	4 (0.14)	0.01
Total mutants (SEQUENOM)		2878 (97.92)	10.10
Total mutants (COSMIC)		2940 (100)	10.32

**Supplementary Table 3 KRAS mutational data from the COSMIC database**

<b>Amino acid change</b>	<b>Nucleotide mutation</b>	<b>Relative mutation distribution in carcinomas, <i>n</i> (%)</b>	<b>Absolute mutation frequency in carcinomas, %</b>
p.G12C	c.34G>T	600 (7.99)	2.84
p.G12S	c.34G>A	448 (5.97)	2.12
p.G12R	c.34G>C	88 (1.17)	0.42
p.G12D	c.35G>A	2551 (33.97)	12.06
p.G12V	c.35G>T	1588 (21.15)	7.51
p.G12A	c.35G>C	493 (6.56)	2.33
p.G13D	c.38G>A	1444 (19.23)	6.83
p.G13A	c.38G>C	7 (0.09)	0.03
p.G13V	c.38G>T	9 (0.12)	0.04
p.G13G	c.39C>A	2 (0.03)	0.01
p.G13G	c.39C>G	1 (0.01)	0
p.G13G	c.39C>T	0 (0)	0
p.A59T	c.175G>A	1 (0.01)	0
p.Q61K	c.181C>A	4 (0.05)	0.05
p.Q61E	c.181C>G	1 (0.01)	0.01
p.Q61L	c.182A>T	11 (0.15)	0.15
p.Q61R	c.182A>G	6 (0.08)	0.08
p.Q61P	c.182A>C	0 (0)	0
p.Q61H	c.183A>C	24 (0.32)	0.32
p.Q61H	c.183A>T	15 (0.20)	0.20
p.A146T	c.436G>A	20 (0.27)	0.09
p.A146P	c.436G>C	2 (0.03)	0.01
Total codon 12		5768 (76.81)	27.28
Total codon 13		1463 (19.48)	6.91
Total codon 61		61 (0.81)	0.81
Total codon 146		22 (0.30)	0.10
Total mutants (SEQUENOM)		7306 (97.28)	34.54

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Total mutants (COSMIC)	7511 (100)	35.51
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**Supplementary Table 4 NRAS mutational data from the COSMIC database**

<b>Amino acid Change</b>	<b>Nucleotide mutation</b>	<b>Relative mutation distribution in carcinomas, <i>n</i> (%)</b>	<b>Absolute mutation frequency in carcinomas, %</b>
p.G12C	c.34G>T	2 (12.50)	0.31
p.G12S	c.34G>A	1 (6.25)	0.16
p.G12R	c.34G>C	0 (0)	0
p.G13R	c.37G>C	1 (6.25)	0.16
p.G13S	c.37G>A	0 (0)	0
p.G13C	c.37G>T	0 (0)	0
p.Q61K	c.181C>A	0 (0)	0
p.Q61E	c.181C>G	0 (0)	0
p.Q61R	c.182A>G	6 (37.50)	0.93
p.Q61P	c.182A>C	0 (0)	0
p.Q61L	c.182A>T	1 (6.25)	0.16
p.Q61H	c.183A>C	0 (0)	0
p.Q61Q	c.183A>G	0 (0)	0
p.Q61H	c.183A>T	0 (0)	0
Total codon 12		3 (18.75)	0.47
Total codon 13		1 (6.25)	0.16
Total codon 61		7 (43.75)	1.09
Total mutants (SEQUENOM)		11 (68.75)	1.71
Total mutants (COSMIC)		16 (100)	2.48

**Supplementary Table 5 *PIKCA3* mutational data from the COSMIC database**

<b>Amino acid change</b>	<b>Nucleotide mutation</b>	<b>Relative mutation distribution in carcinomas, <i>n</i> (%)</b>	<b>Absolute mutation frequency in carcinomas, %</b>
p.G12D	c.35G>A	1 (0.16)	0.02
p.R38H	c.113G>A	3 (0.47)	0.07
p.E81K	c.241G>A	1 (0.16)	0.02
p.R88Q	c.263G>A	6 (0.95)	0.13
p.R93W	c.277C>T	3 (0.47)	0.07
p.G106V	c.317G>T	3 (0.47)	0.07
p.R108H	c.323G>A	4 (0.63)	0.63
p.G118D	c.353G>A	3 (0.47)	0.47
p.P134S	c.400C>T	1 (0.16)	0.02
p.S158L	c.473C>T	1 (0.16)	0.02
p.H160N	c.478C>A	1 (0.16)	0.02
p.K179T	c.536A>C	1 (0.16)	0.02
p.K184E	c.550A>G	1 (0.16)	0.02
p.N345K	c.1035T>A	3 (0.47)	0.07
p.C420R	c.1258T>C	5 (0.79)	0.11
p.P539R	c.1616C>G	1 (0.16)	0.02
p.E542K	c.1624G>A	100 (15.80)	2.17
p.E542Q	c.1624G>C	2 (0.32)	0.04
p.E545K	c.1633G>A	179 (28.28)	3.88
p.Q546K	c.1636C>A	25 (3.95)	0.54
p.Q546E	c.1636C>G	3 (0.47)	0.07
p.H701P	c.2102A>C	0 (0)	0
p.C901F	c.2702G>T	1 (0.16)	0.02
p.M1004I	c.3012G>T	1 (0.16)	0.02
p.G1007R	G3019C	1 (0.16)	0.02
p.H1047Y	c.3139C>T	9 (1.42)	0.20

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p.H1047R	c.3140A>G	121 (19.12)	2.62
p.H1047L	c.3140A>T	22 (3.48)	0.48
p.G1049S	c.3145G>A	0 (0)	0
p.G1049R	c.3145G>C	16 (2.53)	0.35
exon 9		310 (49.43)	6.72
exon 20		170 (26.87)	3.69
Total mutants (SEQUENOM)		515 (81.36)	11.17
Total mutants (COSMIC)		633 (100)	13.73

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