

Table 2. SGX393 single-agent cell-based mutagenesis assays

[SGX393]	Total wells analyzed	Sequenced clones (KD mutant clones)	Mutation(s)	# Occurrences	Frequency by mutation (%)	Fraction of mutated sequences (%)	Residue	Frequency by residue (%)	Fraction of mutated residues (%)
120nM	192	n=48 (21)	Native Bcr-Abl	27	56.3	---	Native Bcr-Abl	56.3	---
			L248R	1	2.1	4.8	248	2.1	4.8
			G250E	2	4.2	9.5	250	6.3	14.3
			G250W	1	2.1	4.8			
			Y253F	3	6.3	14.3			
			Y253H	1	2.1	4.8			
			Y253N	1	2.1	4.8	253	10.4	23.8
			E255K	2	4.2	9.5			
			E255V	2	4.2	9.5			
			F311I	1	2.1	4.8	311	2.1	4.8
			F317C	1	2.1	4.8			
			F317I	2	4.2	9.5	317	14.6	33.3
			F317L	2	4.2	9.5			
			F317V	2	4.2	9.5			
F317V	2	4.2	9.5						
240nM	480	n=243 (204)	Native Bcr-Abl	39	16.0	---	Native Bcr-Abl	16.0	---
			L248M	1	0.4	0.5	248	2.5	2.9
			L248R	4	1.6	2.0			
			L248V	1	0.4	0.5			
			G249D	2	0.8	1.0	249	0.8	1.0
			G250E	47	19.3	23.0			
			Q252E	1	0.4	0.5	250	19.3	23.0
			Q252H	4	1.6	2.0			
			Y253C	3	1.2	1.5	252	2.1	2.5
			Y253F	9	3.7	4.4			
			Y253H	19	7.8	9.3			
			Y253N	6	2.5	2.9			
			E255K	13	5.3	6.4	253	15.2	18.1
			E255V	9	3.7	4.4			
			E258K	4	1.6	2.0			
			F311I	6	2.5	2.9	255	9.1	10.8
			F317C	5	2.1	2.5			
			F317I	9	3.7	4.4	258	1.6	2.0
			F317L	22	9.1	10.8			
			F317S	2	0.8	1.0	311	2.5	2.9
			F317V	34	14.0	16.7			
			N322K	1	0.4	0.5	317	29.6	35.3
			E355G	1	0.4	0.5			
S417Y	1	0.4	0.5						
480nM	288	n=90 (89)	Native Bcr-Abl	1	1.1	---	Native Bcr-Abl	1.1	---
			L248M	1	1.1	1.1	248	6.7	6.7
			L248Q	3	3.3	3.4			
			L248R	2	2.2	2.2			
			G249D	2	2.2	2.2	249	2.2	2.2
			G250E	20	22.2	22.5			
			G250V	4	4.4	4.5	250	26.7	27.0
			Q252H	5	5.6	5.6			
			Y253F	4	4.4	4.5	252	5.6	5.6
			Y253H	6	6.7	6.7			
			Y253N	4	4.4	4.5	253	15.6	15.7
			E255K	1	1.1	1.1			
			E255V	4	4.4	4.5			
			F317C	5	5.6	5.6	255	5.6	5.6
			F317I	3	3.3	3.4			
			F317L	7	7.8	7.9	317	36.7	37.1
			F317V	18	20.0	20.2			
F317V	18	20.0	20.2						
960nM	288	n=56 (56)	Native Bcr-Abl	0	0.0	---	Native Bcr-Abl	0.0	---
			L248M	1	1.8	1.8	248	5.4	5.4
			L248Q	1	1.8	1.8			
			L248R	1	1.8	1.8			
			G250E	8	14.3	14.3	250	14.3	14.3
			Q252H	3	5.4	5.4			
			Y253F	4	7.1	7.1	252	5.4	5.4
			Y253N	2	3.6	3.6			
			E255V	2	3.6	3.6	253	10.7	10.7
			F317C	2	3.6	3.6			
			F317L	3	5.4	5.4	255	3.6	3.6
			F317S	1	1.8	1.8			
			F317V	28	50.0	50.0	317	60.7	60.7
			F317V	28	50.0	50.0			
F317V	28	50.0	50.0						
1920nM	288	n=26 (26)	Native Bcr-Abl	0	0.0	---	Native Bcr-Abl	0.0	---
			L248R	2	7.7	7.7	248	7.7	7.7
			G250E	7	26.9	26.9			
			Y253F	4	15.4	15.4	250	26.9	26.9
			E255V	2	7.7	7.7			
			F317V	11	42.3	42.3	253	15.4	15.4
			F317V	11	42.3	42.3			

NOTE: n values for each SGX393 concentration refer to the total number of sequenced clones. Numbers in parentheses that follow refer to the number of clones carrying mutant Bcr-Abl.