

**Supplementary Table 2. A) Cytosine Methylation Status by Mass Spectrometry.** Cytosine at CpG positions, **2369** and **2383** have high methylation ratio in PC9, H1299 and H460. Cytosine at CpG positions **2331** and **2353** are not. H1975 has C>T mutation at position **2369**, thus no methylation was detected. **B) PC9 mutation frequency by WES after gefitinib treatment;** showing number of new variants detected including SNPs and specifically cytosine based SNPs

**A)**

		Cell Line				
		PC9	PC9	H1299	H1975	H460
<b>CpG Position</b>	<b>EGFR-m_CpG_2331</b>	Neg	Neg	Neg	Neg	Neg
	<b>EGFR-m_CpG_2353</b>	Neg	Neg	Neg	Neg	Neg
	<b>EGFR-m_CpG_2369</b>	0.99	0.89	0.85	Neg	0.87
	<b>EGFR-m_CpG_2383</b>	1	1	1	0.97	1

**B)**

	<b>Variants</b>	<b>SNP</b>	<b>C_SNP</b>
<b>PC9 ctrl vs. Gef</b>	137	72	46
<b>PC9 ctrl vs. Gef</b>	153	98	60