

**Supplementary Table 1: *EZH2* Co-Expression Gene Signature**Top 116 genes co-expressed with *EZH2* across 9 Oncomine studies

Gene Symbol	Description
<b>ALS2CR4</b>	amyotrophic lateral sclerosis 2 (juvenile) chromosome region, candidate 4
<b>ATIC</b>	5-aminoimidazole-4-carboxamide ribonucleotide formyltransferase/IMP cyclohydrolase
<b>ATP2A2</b>	ATPase, Ca++ transporting, cardiac muscle, slow twitch 2
<b>AURKA</b>	aurora kinase A
<b>BIRC5</b>	baculoviral IAP repeat-containing 5
<b>BOP1</b>	block of proliferation 1
<b>BUB1</b>	budding uninhibited by benzimidazoles 1 homolog (yeast)
<b>BUB1B</b>	budding uninhibited by benzimidazoles 1 homolog beta (yeast)
<b>C13orf34</b>	chromosome 13 open reading frame 34
<b>C1orf112</b>	chromosome 1 open reading frame 112
<b>CCNA2</b>	cyclin A2
<b>CCNB1</b>	cyclin B1
<b>CCNB2</b>	cyclin B2
<b>CDC20</b>	cell division cycle 20 homolog ( <i>S. cerevisiae</i> )
<b>CDC25A</b>	cell division cycle 25 homolog A ( <i>S. pombe</i> )
<b>CDC6</b>	cell division cycle 6 homolog ( <i>S. cerevisiae</i> )
<b>CDC7</b>	cell division cycle 7 homolog ( <i>S. cerevisiae</i> )
<b>CDCA3</b>	cell division cycle associated 3
<b>CDCA8</b>	cell division cycle associated 8
<b>CDK1</b>	cyclin-dependent kinase 1
<b>CDKN3</b>	cyclin-dependent kinase inhibitor 3
<b>CENPE</b>	centromere protein E, 312kDa
<b>CENPF</b>	centromere protein F, 350/400kDa (mitosin)
<b>CENPM</b>	centromere protein M
<b>CENPW</b>	centromere protein W
<b>CHEK1</b>	CHK1 checkpoint homolog ( <i>S. pombe</i> )
<b>CKAP2</b>	cytoskeleton associated protein 2
<b>CKS1B</b>	CDC28 protein kinase regulatory subunit 1B
<b>CKS2</b>	CDC28 protein kinase regulatory subunit 2
<b>CSE1L</b>	CSE1 chromosome segregation 1-like (yeast)
<b>CSTF1</b>	cleavage stimulation factor, 3' pre-RNA, subunit 1, 50kDa
<b>CTPS</b>	CTP synthase
<b>DBF4</b>	DBF4 homolog ( <i>S. cerevisiae</i> )
<b>DBN1</b>	drebrin 1
<b>DEPDC1</b>	DEP domain containing 1
<b>DLGAP5</b>	discs, large ( <i>Drosophila</i> ) homolog-associated protein 5
<b>DTL</b>	denticleless homolog ( <i>Drosophila</i> )
<b>ECT2</b>	Epithelial cell transforming sequence 2 oncogene
<b>ESCO2</b>	establishment of cohesion 1 homolog 2 ( <i>S. cerevisiae</i> )

<b>ESPL1</b>	extra spindle pole bodies homolog 1 ( <i>S. cerevisiae</i> )
<b>EXO1</b>	exonuclease 1
<b>EZH2</b>	enhancer of zeste homolog 2 ( <i>Drosophila</i> )
<b>FANCI</b>	Fanconi anemia, complementation group I
<b>FBXO5</b>	F-box protein 5
<b>FEN1</b>	flap structure-specific endonuclease 1
<b>FOXM1</b>	forkhead box M1
<b>GINS1</b>	GINS complex subunit 1 (Psf1 homolog)
<b>GINS2</b>	GINS complex subunit 2 (Psf2 homolog)
<b>GMNN</b>	geminin, DNA replication inhibitor
<b>GOT1</b>	glutamic-oxaloacetic transaminase 1, soluble (aspartate aminotransferase 1)
<b>GPD2</b>	glycerol-3-phosphate dehydrogenase 2 (mitochondrial)
<b>H2AFX</b>	H2A histone family, member X
<b>H2AFZ</b>	H2A histone family, member Z
<b>HMGA1</b>	high mobility group AT-hook 1
<b>HMGB2</b>	high-mobility group box 2
<b>HMMR</b>	hyaluronan-mediated motility receptor (RHAMM)
<b>HN1</b>	hematological and neurological expressed 1
<b>KIAA0101</b>	KIAA0101
<b>KIF11</b>	kinesin family member 11
<b>KIF14</b>	kinesin family member 14
<b>KIF15</b>	kinesin family member 15
<b>KIF18A</b>	kinesin family member 18A
<b>KIF18B</b>	kinesin family member 18B
<b>KIF20A</b>	kinesin family member 20A
<b>KIF20B</b>	kinesin family member 20B
<b>KIF23</b>	kinesin family member 23
<b>KIF2C</b>	kinesin family member 2C
<b>KIFC1</b>	kinesin family member C1
<b>KNTC1</b>	kinetochore associated 1
<b>KPNA2</b>	karyopherin alpha 2 (RAG cohort 1, importin alpha 1)
<b>LMNB1</b>	lamin B1
<b>LMNB2</b>	lamin B2
<b>MAD2L1</b>	MAD2 mitotic arrest deficient-like 1 (yeast)
<b>MCM10</b>	minichromosome maintenance complex component 10
<b>MCM7</b>	minichromosome maintenance complex component 7
<b>MKI67</b>	antigen identified by monoclonal antibody Ki-67
<b>MLF1IP</b>	MLF1 interacting protein
<b>MTHFD2</b>	methylenetetrahydrofolate dehydrogenase (NADP+ dependent) 2,
<b>NCAPG</b>	non-SMC condensin I complex, subunit G
<b>NCAPG2</b>	non-SMC condensin II complex, subunit G2
<b>NCAPH</b>	non-SMC condensin I complex, subunit H
<b>NDC80</b>	NDC80 homolog, kinetochore complex component ( <i>S. cerevisiae</i> )
<b>NUP205</b>	nucleoporin 205kDa
<b>OLA1</b>	Obg-like ATPase 1
<b>PAICS</b>	phosphoribosylaminoimidazole carboxylase

<b>PBK</b>	PDZ binding kinase
<b>PCNA</b>	proliferating cell nuclear antigen
<b>PFN2</b>	profilin 2
<b>PLK4</b>	polo-like kinase 4
<b>POLA1</b>	polymerase (DNA directed), alpha 1, catalytic subunit
<b>POLE2</b>	polymerase (DNA directed), epsilon 2 (p59 subunit)
<b>PPAT</b>	Phosphoribosyl pyrophosphate amidotransferase
<b>PRIM1</b>	primase, DNA, polypeptide 1 (49kDa)
<b>PSMD14</b>	proteasome (prosome, macropain) 26S subunit, non-ATPase, 14
<b>PTTG1</b>	pituitary tumor-transforming 1
<b>RFC4</b>	replication factor C (activator 1) 4, 37kDa
<b>SLC7A5</b>	solute carrier family 7 (cationic amino acid transporter, y+ system), member 5
<b>SMC2</b>	structural maintenance of chromosomes 2
<b>SMC4</b>	structural maintenance of chromosomes 4
<b>SPAG5</b>	sperm associated antigen 5
<b>SRM</b>	spermidine synthase
<b>STIL</b>	SCL/TAL1 interrupting locus
<b>STMN1</b>	stathmin 1
<b>TACC3</b>	transforming, acidic coiled-coil containing protein 3
<b>TFRC</b>	Transferrin receptor (p90, CD71)
<b>TMEM106C</b>	transmembrane protein 106C
<b>TOP2A</b>	Topoisomerase (DNA) II alpha 170kDa
<b>TOPBP1</b>	topoisomerase (DNA) II binding protein 1
<b>TPX2</b>	TPX2, microtubule-associated, homolog (Xenopus laevis)
<b>TRIP13</b>	thyroid hormone receptor interactor 13
<b>TROAP</b>	trophinin associated protein (tastin)
<b>TYMS</b>	thymidylate synthetase
<b>UBE2C</b>	ubiquitin-conjugating enzyme E2C
<b>UBE2S</b>	ubiquitin-conjugating enzyme E2S
<b>UNG</b>	uracil-DNA glycosylase
<b>WDR12</b>	WD repeat domain 12

**Supplementary Table 2: Gene function enrichment analysis of *EZH2* co-expressed gene signature**

C1: number of genes in a cluster or list that have this annotation term

C2: number of annotated genes in this cluster or list

C3: number of all genes on array that have this annotation term

C4: number of all annotated genes on array

P-value: binomial approximated p-value for hypergeometric distribution

**\*\*\*Gene Ontology\*\*\***

C1	C2	C3	C4	P-value	Term Name
81	112	5956	12364	0	protein binding
73	112	3967	12364	0	nucleus
58	112	3873	12364	0.000005	cytoplasm
49	112	439	12364	0	cell cycle
38	112	243	12364	0	cell division
36	112	186	12364	0	mitosis
36	112	1621	12364	0	nucleotide binding
35	112	1250	12364	0	ATP binding
30	112	436	12364	0	nucleoplasm
24	112	667	12364	0	cytoskeleton
20	112	144	12364	0	DNA replication
18	112	294	12364	0	cell proliferation
18	112	103	12364	0	spindle
16	112	203	12364	0	microtubule
15	112	224	12364	0	DNA repair
14	112	155	12364	0	chromosome
12	112	46	12364	0	phosphoinositide-mediated signaling
11	112	53	12364	0	condensed chromosome kinetochore
11	112	71	12364	0	kinetochore
11	112	54	12364	0	microtubule motor activity
11	112	65	12364	0	microtubule-based movement
11	112	109	12364	0	motor activity
10	112	236	12364	0.000062	response to DNA damage stimulus
10	112	47	12364	0	spindle pole
8	112	59	12364	0	anaphase-promoting complex-dependent proteasomal degradation
8	112	41	12364	0	chromosome segregation
7	112	47	12364	0	chromosome, centromeric region
7	112	135	12364	0.000247	microtubule organizing center
7	112	14	12364	0	spindle organization
6	112	12	12364	0	chromosome condensation

6	112	58	12364	0.000017	negative regulation of mitotic ubiquitin-protein ligase
6	112	61	12364	0.000022	positive regulation of mitotic ubiquitin-protein ligase
6	112	38	12364	0.000002	regulation of cyclin-dependent protein kinase activity
5	112	49	12364	0.000092	microtubule cytoskeleton
5	112	14	12364	0	mitotic sister chromatid segregation
5	112	25	12364	0.000004	spindle microtubule
4	112	19	12364	0.00003	chromosome organization
4	112	23	12364	0.000063	condensed chromosome
4	112	5	12364	0	condensin complex
4	112	36	12364	0.000347	DNA metabolic process
4	112	44	12364	0.000734	G1/S transition of mitotic cell cycle
4	112	8	12364	0.000001	M phase of mitotic cell cycle
4	112	19	12364	0.00003	midbody
4	112	12	12364	0.000005	mitotic cell cycle checkpoint
4	112	11	12364	0.000004	mitotic cell cycle spindle assembly checkpoint
4	112	12	12364	0.000005	mitotic chromosome condensation
4	112	26	12364	0.000101	nuclear chromosome
3	112	6	12364	0.000025	'de novo' IMP biosynthetic process
3	112	20	12364	0.000846	anaphase-promoting complex
3	112	14	12364	0.000302	DNA unwinding involved in replication
3	112	20	12364	0.000846	G2/M transition of mitotic cell cycle
3	112	13	12364	0.000243	kinesin complex
3	112	15	12364	0.000369	male germ cell nucleus
3	112	10	12364	0.000113	mitotic cell cycle
3	112	15	12364	0.000369	mitotic spindle organization
3	112	17	12364	0.00053	nucleotide-excision repair, DNA gap filling
3	112	13	12364	0.000243	oocyte maturation
3	112	8	12364	0.000059	outer kinetochore of condensed chromosome
3	112	15	12364	0.000369	purine nucleotide biosynthetic process
3	112	15	12364	0.000369	regulation of mitosis
3	112	11	12364	0.000149	replication fork
2	112	5	12364	0.000987	lamin filament
2	112	5	12364	0.000987	purine base biosynthetic process
2	112	3	12364	0.00036	regulation of chromosome segregation

64 reported significant, 0 expected false positive

(209 terms assessed for enrichment at p-value threshold 0.001)

### Supplementary Table 3: Chou Talalay Combination Index Analysis

H23 DZNep + Etoposide CI Simulations					SW1573 DZNep + Etoposide CI Simulations					H460 DZNep + Etoposide CI Simulations				
Fa	CI	Est. s.d.	Etop	DZNep	Fa	CI	Est. s.d.	Etop	DZNep	Fa	CI	Est. s.d.	Etop	DZNep
0.02	1E+12	6E+14	0.002	0.002	0.02	103.7	187.1	0.003	0.003	0.02	19.85	45.27	0.001	0.001
0.05	2E+08	2E+10	0.006	0.006	0.05	11.34	11.85	0.012	0.012	0.05	10.29	17.48	0.004	0.004
0.1	3.2E+05	6.2E+06	0.020	0.020	0.1	3.154	2.540	0.036	0.036	0.1	6.175	8.073	0.014	0.014
0.15	5586	40440	0.040	0.040	0.15	1.996	1.617	0.069	0.069	0.15	4.510	4.924	0.030	0.030
0.2	260.4	908.6	0.067	0.067	0.2	1.607	1.244	0.114	0.114	0.2	3.562	3.355	0.055	0.055
0.25	20.85	38.69	0.104	0.104	0.25	1.412	1.012	0.171	0.171	0.25	2.932	2.422	0.090	0.090
0.3	2.449	2.339	0.152	0.152	0.3	1.287	0.845	0.244	0.244	0.3	2.474	1.807	0.139	0.139
0.35	0.512	0.190	0.215	0.215	0.35	1.196	0.717	0.337	0.337	0.35	2.121	1.376	0.206	0.206
0.4	0.280	0.058	0.297	0.297	0.4	1.122	0.614	0.456	0.456	0.4	1.837	1.061	0.297	0.297
0.45	0.268	0.050	0.405	0.405	0.45	1.058	0.530	0.610	0.610	0.45	1.600	0.824	0.422	0.422
0.5	0.291	0.050	0.549	0.549	0.5	1.001	0.461	0.810	0.810	0.5	1.398	0.643	0.596	0.596
0.55	0.323	0.052	0.744	0.744	0.55	0.948	0.405	1.077	1.077	0.55	1.221	0.504	0.842	0.842
0.6	0.360	0.054	1.014	1.014	0.6	0.898	0.360	1.439	1.439	0.6	1.064	0.400	1.197	1.197
0.65	0.404	0.059	1.400	1.400	0.65	0.849	0.326	1.948	1.948	0.65	0.921	0.326	1.728	1.728
0.7	0.456	0.065	1.978	1.978	0.7	0.799	0.304	2.693	2.693	0.7	0.790	0.276	2.559	2.559
0.75	0.521	0.077	2.893	2.893	0.75	0.748	0.294	3.845	3.845	0.75	0.667	0.246	3.943	3.943
0.8	0.608	0.096	4.470	4.470	0.8	0.694	0.296	5.781	5.781	0.8	0.550	0.230	6.467	6.467
0.85	0.732	0.131	7.571	7.571	0.85	0.634	0.311	9.471	9.471	0.85	0.435	0.221	11.77	11.77
0.9	0.937	0.202	15.24	15.24	0.9	0.562	0.339	18.24	18.24	0.9	0.319	0.209	26.08	26.08
0.95	1.396	0.400	47.20	47.20	0.95	0.463	0.384	52.62	52.62	0.95	0.193	0.181	94.31	94.31
0.99	3.370	1.587	573.2	573.2	0.99	0.301	0.450	546.2	546.2	0.99	0.064	0.112	1612	1612
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H23 GSK126 + Etoposide CI Simulations					SW1573 GSK126 + Etoposide CI Simulations					H460 GSK126 + Etoposide CI Simulations				
Fa	CI	Est. s.d.	Etop	GSK126	Fa	CI	Est. s.d.	Etop	GSK126	Fa	CI	Est. s.d.	Etop	GSK126
0.02	0.743	0.307	0.024	0.024	0.02	55.27	406.2	0.006	0.006	0.02	76.45	168.7	0.003	0.003
0.05	0.813	0.254	0.073	0.073	0.05	21.45	103.9	0.020	0.020	0.05	30.72	49.90	0.011	0.011
0.1	0.895	0.217	0.175	0.175	0.1	10.73	35.36	0.048	0.048	0.1	14.97	18.65	0.033	0.033
0.15	0.958	0.197	0.302	0.302	0.15	7.059	18.05	0.084	0.084	0.15	9.593	9.982	0.065	0.065
0.2	1.011	0.187	0.456	0.456	0.2	5.164	10.83	0.128	0.128	0.2	6.861	6.168	0.107	0.107
0.25	1.060	0.183	0.640	0.640	0.25	3.993	7.080	0.180	0.180	0.25	5.202	4.109	0.161	0.161
0.3	1.105	0.185	0.861	0.861	0.3	3.192	4.861	0.244	0.244	0.3	4.085	2.862	0.230	0.230
0.35	1.149	0.192	1.126	1.126	0.35	2.605	3.439	0.320	0.320	0.35	3.280	2.049	0.319	0.319
0.4	1.193	0.204	1.449	1.449	0.4	2.154	2.477	0.414	0.414	0.4	2.671	1.492	0.433	0.433
0.45	1.237	0.219	1.844	1.844	0.45	1.796	1.799	0.529	0.529	0.45	2.193	1.099	0.580	0.580
0.5	1.282	0.240	2.337	2.337	0.5	1.503	1.308	0.674	0.674	0.5	1.808	0.815	0.773	0.773
0.55	1.329	0.265	2.960	2.960	0.55	1.258	0.947	0.857	0.857	0.55	1.491	0.608	1.029	1.029
0.6	1.380	0.295	3.768	3.768	0.6	1.049	0.680	1.096	1.096	0.6	1.224	0.459	1.379	1.379
0.65	1.435	0.331	4.847	4.847	0.65	0.868	0.483	1.416	1.416	0.65	0.997	0.353	1.872	1.872
0.7	1.498	0.375	6.343	6.343	0.7	0.709	0.343	1.863	1.863	0.7	0.801	0.281	2.593	2.593
0.75	1.570	0.430	8.530	8.530	0.75	0.567	0.252	2.519	2.519	0.75	0.629	0.231	3.714	3.714
0.8	1.658	0.502	11.97	11.97	0.8	0.440	0.201	3.559	3.559	0.8	0.477	0.197	5.603	5.603
0.85	1.772	0.602	18.05	18.05	0.85	0.323	0.179	5.407	5.407	0.85	0.342	0.170	9.217	9.217
0.9	1.937	0.758	31.14	31.14	0.9	0.214	0.165	9.423	9.423	0.9	0.219	0.142	17.85	17.85
0.95	2.240	1.076	75.14	75.14	0.95	0.110	0.142	23.11	23.11	0.95	0.107	0.116	51.94	51.94
0.99	3.101	2.159	525.8	525.8	0.99	0.026	0.087	167.7	167.7	0.99	0.023	0.223	549.5	549.5

# Supplementary Table 3 Continued

HCC15 DZNep + Etoposide CI Simulations					A549 DZNep + Etoposide CI Simulations					H157 DZNep + Etoposide CI Simulations				
Fa	CI	Est. s.d.	Etop	DZNep	Fa	CI	Est. s.d.	Etop	DZNep	Fa	CI	Est. s.d.	Etop	DZNep
0.02	164.6	156.3	0.002	0.002	0.02	159.85	561.11	8E-05	8E-05	0.02	4.250	4.229	0.047	0.047
0.05	14.29	9.971	0.009	0.009	0.05	8.957	16.333	0.001	0.001	0.05	1.211	0.617	0.116	0.116
0.1	2.199	1.132	0.029	0.029	0.1	1.085	1.005	0.005	0.005	0.1	0.642	0.200	0.240	0.240
0.15	0.781	0.324	0.062	0.062	0.15	0.428	0.240	0.014	0.014	0.15	0.523	0.144	0.375	0.375
0.2	0.423	0.152	0.111	0.111	0.2	0.303	0.142	0.031	0.031	0.2	0.483	0.124	0.525	0.525
0.25	0.298	0.099	0.177	0.177	0.25	0.274	0.117	0.062	0.062	0.25	0.467	0.112	0.694	0.694
0.3	0.248	0.077	0.268	0.268	0.3	0.271	0.103	0.111	0.111	0.3	0.461	0.104	0.885	0.885
0.35	0.227	0.067	0.389	0.389	0.35	0.277	0.093	0.191	0.191	0.35	0.460	0.098	1.104	1.104
0.4	0.218	0.061	0.553	0.553	0.4	0.287	0.086	0.316	0.316	0.4	0.461	0.094	1.357	1.357
0.45	0.216	0.059	0.773	0.773	0.45	0.298	0.080	0.513	0.513	0.45	0.465	0.091	1.655	1.655
0.5	0.217	0.059	1.074	1.074	0.5	0.311	0.078	0.823	0.823	0.5	0.469	0.090	2.009	2.009
0.55	0.220	0.060	1.493	1.493	0.55	0.324	0.079	1.321	1.321	0.55	0.474	0.089	2.440	2.440
0.6	0.225	0.064	2.089	2.089	0.6	0.339	0.084	2.142	2.142	0.6	0.480	0.091	2.974	2.974
0.65	0.231	0.070	2.966	2.966	0.65	0.355	0.096	3.546	3.546	0.65	0.487	0.094	3.657	3.657
0.7	0.238	0.079	4.313	4.313	0.7	0.373	0.114	6.077	6.077	0.7	0.494	0.099	4.560	4.560
0.75	0.246	0.090	6.514	6.514	0.75	0.394	0.140	11.00	11.00	0.75	0.503	0.107	5.816	5.816
0.8	0.257	0.106	10.44	10.44	0.8	0.420	0.176	21.68	21.68	0.8	0.514	0.119	7.682	7.682
0.85	0.270	0.128	18.49	18.49	0.85	0.454	0.231	49.32	49.32	0.85	0.527	0.136	10.76	10.76
0.9	0.288	0.164	39.49	39.49	0.9	0.502	0.320	147.0	147.0	0.9	0.546	0.163	16.83	16.83
0.95	0.321	0.235	134.5	134.5	0.95	0.592	0.512	857.0	857.0	0.95	0.578	0.215	34.69	34.69
0.99	0.406	0.469	2017	2017	0.99	0.851	1.245	42135	42135	0.99	0.655	0.367	171.3	171.3
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HCC15 GSK + Etoposide CI Simulations					A549 GSK126 + Etoposide CI Simulations					H157 GSK126 + Etoposide CI Simulations				
Fa	CI	Est. s.d.	Etop	GSK126	Fa	CI	Est. s.d.	Etop	GSK126	Fa	CI	Est. s.d.	Etop	GSK126
0.02	401.6	1637	0.012	0.012	0.02	1.072	1.193	0.001	0.001	0.02	3.010	1.435	0.354	0.354
0.05	3.539	3.885	0.044	0.044	0.05	0.902	0.747	0.004	0.004	0.05	2.040	0.806	0.565	0.565
0.1	0.698	0.428	0.122	0.122	0.1	0.787	0.495	0.019	0.019	0.1	1.502	0.505	0.816	0.816
0.15	0.606	0.319	0.229	0.229	0.15	0.724	0.372	0.047	0.047	0.15	1.243	0.377	1.026	1.026
0.2	0.573	0.264	0.367	0.367	0.2	0.679	0.295	0.093	0.093	0.2	1.078	0.304	1.219	1.219
0.25	0.549	0.228	0.542	0.542	0.25	0.645	0.240	0.163	0.163	0.25	0.959	0.255	1.405	1.405
0.3	0.530	0.202	0.763	0.763	0.3	0.616	0.198	0.266	0.266	0.3	0.865	0.219	1.590	1.590
0.35	0.513	0.185	1.039	1.039	0.35	0.591	0.166	0.417	0.417	0.35	0.788	0.193	1.780	1.780
0.4	0.498	0.174	1.388	1.388	0.4	0.568	0.141	0.633	0.633	0.4	0.723	0.172	1.978	1.978
0.45	0.484	0.168	1.833	1.833	0.45	0.548	0.123	0.945	0.945	0.45	0.665	0.155	2.189	2.189
0.5	0.471	0.166	2.406	2.406	0.5	0.528	0.111	1.399	1.399	0.5	0.613	0.141	2.417	2.417
0.55	0.458	0.167	3.159	3.159	0.55	0.509	0.105	2.072	2.072	0.55	0.565	0.129	2.668	2.668
0.6	0.445	0.172	4.170	4.170	0.6	0.491	0.105	3.094	3.094	0.6	0.519	0.119	2.952	2.952
0.65	0.431	0.179	5.571	5.571	0.65	0.472	0.110	4.699	4.699	0.65	0.476	0.110	3.281	3.281
0.7	0.418	0.188	7.592	7.592	0.7	0.453	0.119	7.345	7.345	0.7	0.434	0.103	3.672	3.672
0.75	0.403	0.200	10.68	10.68	0.75	0.433	0.132	12.01	12.01	0.75	0.392	0.096	4.158	4.158
0.8	0.387	0.215	15.77	15.77	0.8	0.412	0.149	21.09	21.09	0.8	0.349	0.090	4.792	4.792
0.85	0.369	0.234	25.29	25.29	0.85	0.387	0.168	41.71	41.71	0.85	0.303	0.084	5.692	5.692
0.9	0.346	0.259	47.37	47.37	0.9	0.356	0.193	103.1	103.1	0.9	0.251	0.077	7.153	7.153
0.95	0.311	0.297	130.5	130.5	0.95	0.313	0.228	445.2	445.2	0.95	0.185	0.068	10.35	10.35
0.99	0.247	0.369	1224	1224	0.99	0.238	0.286	11262	11262	0.99	0.095	0.049	23.38	23.38

## Supplementary Table 3 Continued

PC9 DZNep + Etoposide CI Simulations				
Fa	CI	Est. s.d.	Etop (μM)	DZNep (μM)
0.02	28.21	24.02	0.040	0.040
0.05	3.420	2.107	0.087	0.087
0.1	0.715	0.328	0.161	0.161
0.15	0.325	0.129	0.237	0.237
0.2	0.219	0.083	0.316	0.316
0.25	0.182	0.069	0.401	0.401
0.3	0.171	0.066	0.495	0.495
0.35	0.170	0.066	0.598	0.598
0.4	0.175	0.068	0.714	0.714
0.45	0.183	0.071	0.846	0.846
0.5	0.192	0.075	0.999	0.999
0.55	0.204	0.079	1.180	1.180
0.6	0.217	0.085	1.399	1.399
0.65	0.233	0.092	1.670	1.670
0.7	0.251	0.101	2.019	2.019
0.75	0.273	0.112	2.487	2.487
0.8	0.301	0.128	3.158	3.158
0.85	0.338	0.150	4.217	4.217
0.9	0.396	0.189	6.190	6.190
0.95	0.510	0.277	11.51	11.51
0.99	0.892	0.649	45.30	45.302
PC9 GSK126 + Etoposide CI Simulations				
Fa	CI	Est. s.d.	Etop (μM)	GSK126 (μM)
0.02	38.43	44.77	0.035	0.035
0.05	5.103	3.771	0.093	0.093
0.1	1.124	0.534	0.202	0.202
0.15	0.519	0.194	0.327	0.327
0.2	0.352	0.123	0.470	0.470
0.25	0.300	0.106	0.633	0.633
0.3	0.290	0.104	0.822	0.822
0.35	0.300	0.110	1.043	1.043
0.4	0.320	0.119	1.302	1.302
0.45	0.348	0.131	1.610	1.610
0.5	0.382	0.146	1.984	1.984
0.55	0.422	0.164	2.444	2.444
0.6	0.470	0.187	3.024	3.024
0.65	0.526	0.215	3.775	3.775
0.7	0.595	0.252	4.786	4.786
0.75	0.682	0.302	6.214	6.214
0.8	0.798	0.373	8.380	8.380
0.85	0.966	0.485	12.04	12.04
0.9	1.244	0.691	19.47	19.47
0.95	1.874	1.222	42.32	42.32
0.99	4.632	4.204	235.3	235.3

Supplementary Table 4: Mutational Data of NSCLC Cell Lines Used in this Study

Cell Line	Gender	Age	Ethnicity	Subtype	Stage	Smoke (pack/yr)	Biopsy	PIK3CA	Kras	Nras	p53	SMARCA4	EGFR	CDKN2A	LKB1
H1975	F	N/A	N/A	AdC	N/A	non-smoker	primary	G118D	WT	WT	R273H	WT	T790M, L858R	E69*	Q37* het
H2030	M	N/A	N/A	AdC	N/A	unknown	metastasis	WT	G12C	WT	G262V	deletion	WT	WT	E317*
HCC4006	M	N/A	Caucasian	AdC	N/A	non-smoker	N/A	WT	WT	WT	N/A	N/A	L747_E749del	N/A	WT
A549	M	58	Caucasian	AdC	N/A	unknown	primary	WT	G12S	WT	WT	Q729*	WT	WT	deletion Q37*
HCC2450	M	52	Caucasian	SCC	IIIB	unknown	primary	H1047R	N/A	N/A	deletion	N/A	N/A	N/A	N/A
PC9	M	45	East Asian	AdC	N/A	unknown	N/A	WT	WT	WT	N/A	N/A	E746_A750del	G67V	WT
Calu1	M	47	Caucasian	EpiC	III	unknown	pleura	WT	G12C	WT	WT	WT	WT	WT	WT
H1650	M	27	Caucasian	AdC (BAC)	IIIB	10	pleura	WT	WT	WT	deletion	WT	E746X	WT	WT
H522	M	60	Caucasian	AdC	II	60	primary	WT	WT	WT	P191*	P270*	WT	WT	WT
H2126	M	65	Caucasian	LC	IIIB	non-smoker	pleura	WT	WT	WT	E62*	W764R	WT	deletion	deletion
H157	M	59	Caucasian	SCC	IIIB	50	metastasis	WT	G12R	WT	E298*	T58*	WT	E69*	deletion
H1299	M	43	Caucasian	LC	IIIA	50	lymph node	WT	WT	WT	WT	Y560*	WT	WT	hypermeth
HCC15	M	47	Afr-American	SCC	N/A	unknown	primary	WT	WT	mut	WT	M272*	WT	WT	WT
HCC827	F	39	Caucasian	AdC	N/A	4	N/A	WT	WT	WT	V218del	WT	E746_A750del	N/A	WT
H322	M	52	Caucasian	AdC (BAC)	IV	60	lymph node	WT	WT	WT	N/A	N/A	WT	WT	WT
H2009	F	68	Caucasian	AdC	IV	30	lymph node	E711D	G12A	WT	R273L	WT	WT	WT	WT
Sw1573	F	44	Caucasian	AdC	N/A	unknown	primary	K111E	G12C	WT	WT	WT	WT	WT	WT
Calu6	F	61	Caucasian	AdC (Ana)	N/A	unknown	primary	N/A	WT	WT	R196*	WT	WT	WT	WT
H441	M	33	N/A	AdC (pap)	IIIA	non-smoker	pleura	WT	G12V	WT	R158L	WT	WT	WT	WT
HCC95	M	64	Caucasian	SCC	IV	unknown	pleura	Amplified	WT	WT	WT	WT	WT	WT	WT
H520	M	N/A	N/A	SCC	N/A	non-smoker	primary	WT	WT	WT	WT	WT	WT	G45*	WT
H460	M	N/A	N/A	LC	N/A	non-smoker	primary	E545K	WT	WT	WT	WT	WT	WT	WT
Calu3	M	25	Caucasian	AdC	N/A	unknown	pleura	Amplified	WT	WT	M237I	WT	WT	WT	WT
											Q16L, Q16L,				
H2122	F	46	Caucasian	AdC	IV	30	pleura	WT	G12C	WT	C176F	WT	WT	WT	WT
H23	M	51	Afr-American	AdC	N/A	40	primary	WT	G12C	WT	M246I	K1533N	WT	WT	WT
H3255	F	47	Caucasian	AdC	IIIB	non-smoker	N/A	Amplified	G12C	WT	N/A	N/A	L858R	N/A	N/A
Cell Line	Gender	Age	Ethnicity	Subtype	Stage	Smoke (pack/yr)	Biopsy	PIK3CA	Kras	Nras	p53	SMARCA4	EGFR	CDKN2A	LKB1

**Supplementary Table 5: Shared differentially expressed genes in *EGFR* and *BRG1* mutant NSCLC cell lines**

Probe	Symbol	Description	p-value	log2 fold change for Brg1mut - WT	log2 fold change for EGFRmut - WT
201983_s_at	<b>EGFR</b>	epidermal growth factor receptor	0.014	2.178	2.232
210986_s_at	<b>TPM1</b>	tropomyosin 1 (alpha)	0.007	1.525	1.743
218368_s_at	<b>TNFRSF12A</b>	tumor necrosis factor receptor, 12A	0.004	1.172	1.173
205130_at	<b>MOK</b>	MOK protein kinase	0.013	0.928	0.891
217785_s_at	<b>YKT6</b>	YKT6 v-SNARE homolog (S. cerevisiae)	0.01	0.913	1.196
219088_s_at	<b>ZNF576</b>	zinc finger protein 576	7E-04	0.907	1.008
202408_s_at	<b>PRPF31</b>	PRP31 pre-mRNA processing factor 31	0.001	0.755	0.974
218261_at	<b>AP1M2</b>	adaptor-related protein complex 1, mu	0.013	0.748	0.937
202578_s_at	<b>DDX19A</b>	DEAD (Asp-Glu-Ala-Asp) box 19A	0.002	0.716	0.531
201144_s_at	<b>EIF2S1</b>	eukaryotic translation initiation factor 2	0.012	0.69	0.79
216299_s_at	<b>XRCC3</b>	X-ray repair complementing defective	0.003	0.574	0.572
217784_at	<b>YKT6</b>	YKT6 v-SNARE homolog (S. cerevisiae)	0.002	0.531	0.81
1773_at	<b>FNTB</b>	farnesyltransferase, CAAX box, beta	0.003	0.514	0.408
218408_at	<b>TIMM10</b>	membrane 10 homolog (yeast)	0.004	0.475	0.727
221769_at	<b>SPSB3</b>	SOCS box containing 3	0.008	0.471	0.614
202927_at	<b>PIN1</b>	peptidylprolyl cis/ NIMA-interacting 1	0.007	0.456	0.451
211289_x_at	<b>CDK11</b>	cyclin dependent kinase 11	0.006	0.438	0.373
218206_x_at	<b>SCAND1</b>	SCAN domain containing 1	0.005	0.396	0.543
214244_s_at	<b>ATP6V0E1</b>	ATPase V0 subunit e1	0.005	0.384	0.481
221907_at	<b>TRMT61A</b>	tRNA methyltransferase	0.006	0.378	0.522
221650_s_at	<b>MED18</b>	mediator complex subunit 18	0.008	0.251	0.29