

# **Supporting Information**

## **Dual inhibition of EGFR at protein and activity level via combinatorial blocking of PI4KII $\alpha$ as anti-tumor strategy**

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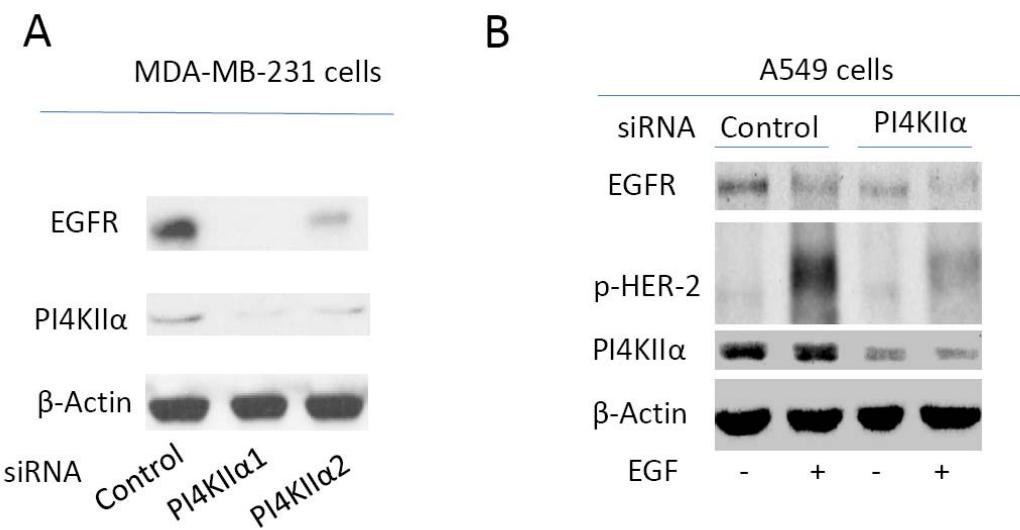
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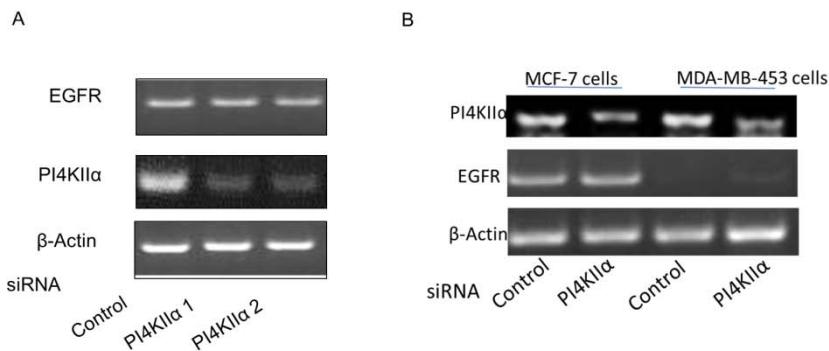
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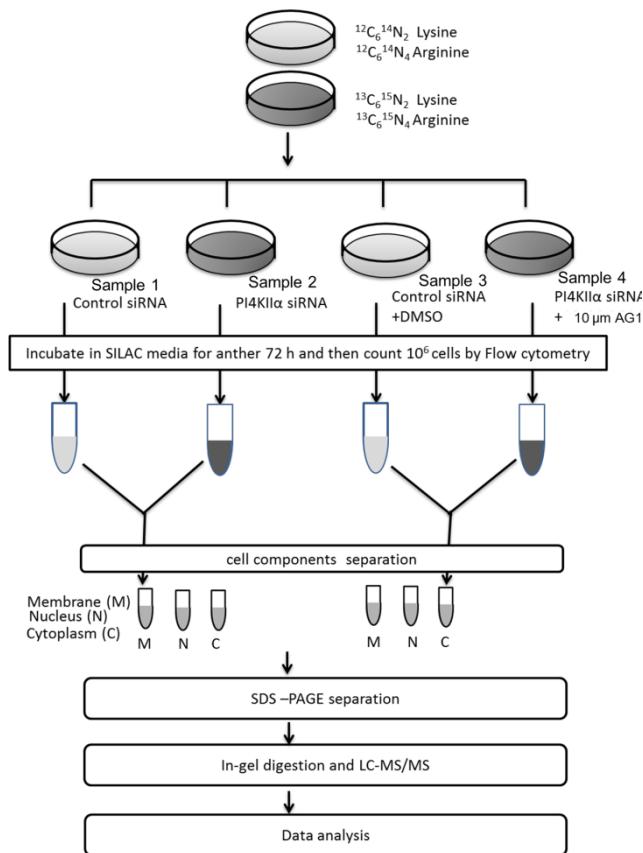
**Running Title:** PI4KII $\alpha$ , an ideal combinatorial target for EGFR treatment



**Figure S1. Effect of PI4KII $\alpha$  knockdown in different cell lines on EGFR protein levels.** **A)** MDA-MB-231 cells. **B)** A549 cells. Cells were transfected with either control siRNA or PI4KII $\alpha$  siRNA (PI4KII $\alpha$  1 and PI4KII $\alpha$  2 refer to siRNA targeting different site of the PI4KII $\alpha$  mRNA) for 72 hours, and then treated 100 ng/ml EGF for 10 minutes, except controls, and proteins indicated measured by Western blot. All results presented above represent data from three independent experiments.



**Figure S2. PI4KII $\alpha$  knockdown not affect EGFR transcription level.** **A)** MCF-7 cells were transfected with control siRNA or two different PI4KII $\alpha$  siRNAs for 72 hours. mRNA levels of EGFR and PI4KII $\alpha$ . **B)** MCF-7 cells and MDA-MB-435 cells were transfected with control or PI4KII $\alpha$  siRNA for 72 hours, and levels of EGFR and PI4KII $\alpha$  mRNA measured.



**Figure S3. Proteomics analysis of molecular networks affected by PI4KIIα knockdown or dual inhibition by SILAC and LC-MS/MS.** Schematic flowchart for the discovery of molecular networks regulated by PI4K and EGFR dual inhibition in MCF-7 cells. Detailed information was described in method section.

**Table S1. Protein targets identified by SILAC and LC-MS/MS for control RNAi MCF-7 cells (sample 1) and PI4KII $\alpha$  RNAi MCF-7 cells (sample 2).**

Accession	Description	Heavy/Light*
<b>O94776</b>	Metastasis-associated protein MTA2	0.22
<b>P04040</b>	Catalase	0.24
<b>P14923</b>	Junction plakoglobin	0.26
<b>Q08945</b>	FACT complex subunit SSRP1	0.26
<b>Q9UMS4</b>	Pre-mRNA-processing factor 19	0.26
<b>P12270</b>	Nucleoprotein TPR	0.26
<b>P08670</b>	Vimentin	0.27
<b>P09874</b>	Poly [ADP-ribose] polymerase 1	0.28
<b>Q9BTV4</b>	Transmembrane protein 43	0.29
<b>Q14690</b>	Protein RRP5 homolog	0.30
<b>Q15269</b>	Periodic tryptophan protein 2 homolog	0.31
<b>P14866</b>	Heterogeneous nuclear ribonucleoprotein L	0.32
<b>P23246</b>	Splicing factor, proline- and glutamine-rich	0.32
<b>Q9BQ39</b>	ATP-dependent RNA helicase DDX50	0.33
<b>O43143</b>	Putative pre-mRNA-splicing factor ATP-dependent RNA helicase DHX15	0.34
<b>Q13151</b>	Heterogeneous nuclear ribonucleoprotein A0	0.34
<b>O75533</b>	Splicing factor 3B subunit 1	0.35
<b>H0Y8X1</b>	Succinate dehydrogenase [ubiquinone] flavoprotein subunit, mitochondrial	0.35
<b>H0YDK2</b>	Histone-binding protein RBBP4 (Fragment)	0.35
<b>Q99436</b>	Proteasome subunit beta type-7	0.35
<b>P36871</b>	Phosphoglucomutase-1	0.36
<b>Q9NVP1</b>	ATP-dependent RNA helicase DDX18	0.36
<b>P35237</b>	Serpin B6	0.37
<b>Q13347</b>	Eukaryotic translation initiation factor 3 subunit I	0.38
<b>P30740</b>	Leukocyte elastase inhibitor	0.38
<b>P48147</b>	Prolyl endopeptidase	0.39
<b>E9PBF6</b>	Lamin-B1	0.39

<b>Q08211</b>	ATP-dependent RNA helicase A	0.39
<b>P12004</b>	Proliferating cell nuclear antigen	0.40
<b>P04080</b>	Cystatin-B	0.40
<b>Q96GQ7</b>	Probable ATP-dependent RNA helicase DDX27	0.40
<b>P14550</b>	Alcohol dehydrogenase [NADP(+)]	0.40
<b>Q16795</b>	NADH dehydrogenase [ubiquinone] 1 alpha subcomplex subunit 9, mitochondrial	0.41
<b>P24539</b>	ATP synthase subunit b, mitochondrial	0.42
<b>O15427</b>	Monocarboxylate transporter 4	0.42
<b>Q8TDN6</b>	Ribosome biogenesis protein BRX1 homolog	0.42
<b>P32119</b>	Peroxiredoxin-2	0.43
<b>P22087</b>	rRNA 2'-O-methyltransferase fibrillarin	0.43
<b>P05386</b>	60S acidic ribosomal protein P1	0.43
<b>Q15645</b>	Pachytene checkpoint protein 2 homolog	0.43
<b>O00487</b>	26S proteasome non-ATPase regulatory subunit 14	0.44
<b>Q6S8J3</b>	POTE ankyrin domain family member E	0.44
<b>Q09666</b>	Neuroblast differentiation-associated protein AHNAK	0.45
<b>P16152</b>	Carbonyl reductase [NADPH] 1	0.45
<b>P28072</b>	Proteasome subunit beta type-6	0.45
<b>Q7KZF4</b>	Staphylococcal nuclease domain-containing protein 1	0.45
<b>P52907</b>	F-actin-capping protein subunit alpha-1	0.45
<b>P52292</b>	Importin subunit alpha-2	0.45
<b>P25787</b>	Proteasome subunit alpha type-2	0.46
<b>P08574</b>	Cytochrome c1, heme protein, mitochondrial	0.46
<b>P24534</b>	Elongation factor 1-beta	0.46
<b>Q15393</b>	Splicing factor 3B subunit 3	0.46
<b>P28066</b>	Proteasome subunit alpha type-5	0.47
<b>P26447</b>	Protein S100-A4	0.47
<b>P37802</b>	Transgelin-2	0.47
<b>O75874</b>	Isocitrate dehydrogenase [NADP] cytoplasmic	0.47
<b>P20618</b>	Proteasome subunit beta type-1	0.48
<b>Q9Y2X3</b>	Nucleolar protein 58	0.48

P62269	40S ribosomal protein S18	0.48
O95678	Keratin, type II cytoskeletal 75	0.48
P80723	Brain acid soluble protein 1	0.48
Q14566	DNA replication licensing factor MCM6	0.48
P00966	Argininosuccinate synthase	0.48
P22314	Ubiquitin-like modifier-activating enzyme 1	0.48
P28070	Proteasome subunit beta type-4	0.48
O75643	U5 small nuclear ribonucleoprotein 200 kDa helicase	0.49
P13010	X-ray repair cross-complementing protein 5	0.49
Q6P2Q9	Pre-mRNA-processing-splicing factor 8	0.49
P49915	GMP synthase [glutamine-hydrolyzing]	0.49
P30041	Peroxiredoxin-6	0.49
E9PBV3	Suprabasin	0.50
P53985	Monocarboxylate transporter 1	0.50
Q99497	Protein DJ-1	0.50
P34932	Heat shock 70 kDa protein 4	0.50
P41091	Eukaryotic translation initiation factor 2 subunit 3	0.50
Q96CS3	FAS-associated factor 2	0.50
P78527	DNA-dependent protein kinase catalytic subunit	0.51
Q9NQR4	Omega-amidase NIT2	0.51
Q9BYZ2	L-lactate dehydrogenase A-like 6B	0.51
O00116	Alkyldihydroxyacetonephosphate synthase, peroxisomal	0.51
O43242	26S proteasome non-ATPase regulatory subunit 3	0.51
P08238	Heat shock protein HSP 90-beta	0.51
P49257	Protein ERGIC-53	0.52
O95373	Importin-7	0.52
P31930	Cytochrome b-c1 complex subunit 1, mitochondrial	0.52
P30084	Enoyl-CoA hydratase, mitochondrial	0.53
P61353	60S ribosomal protein L27	0.53
P17931	Galectin-3	0.53
P25705	ATP synthase subunit alpha, mitochondrial	0.53

P52597	Heterogeneous nuclear ribonucleoprotein F	0.53
Q9BWD1	Acetyl-CoA acetyltransferase, cytosolic	0.53
P62081	40S ribosomal protein S7	0.53
P25398	40S ribosomal protein S12	0.53
P60903	Protein S100-A10	0.54
P62854	40S ribosomal protein S26	0.54
P48637	Glutathione synthetase	0.54
P46777	60S ribosomal protein L5	0.54
E9PK82	40S ribosomal protein S3 (Fragment)	0.54
O14980	Exportin-1	0.54
P43121	Cell surface glycoprotein MUC18	0.54
O43707	Alpha-actinin-4	0.54
Q14126	Desmoglein-2	0.54
Q9BVC6	Transmembrane protein 109	0.55
O75131	Copine-3	0.55
P08754	Guanine nucleotide-binding protein G(k) subunit alpha	0.55
P23526	Adenosylhomocysteinase	0.55
P62906	60S ribosomal protein L10a	0.55
P10599	Thioredoxin	0.55
P49721	Proteasome subunit beta type-2	0.56
P37837	Transaldolase	0.56
Q6ZS74	Ras-related protein Ral-B	0.56
P00491	Purine nucleoside phosphorylase	0.56
P40429	60S ribosomal protein L13a	0.56
P30050	60S ribosomal protein L12	0.57
Q08380	Galectin-3-binding protein	0.57
P26373	60S ribosomal protein L13	0.57
P27348	14-3-3 protein theta	0.57
P17301	Integrin alpha-2	0.58
P62701	40S ribosomal protein S4, X isoform	0.58
P31947	14-3-3 protein sigma	0.58

P15121	Aldose reductase	0.59
P08708	40S ribosomal protein S17	0.59
P62266	40S ribosomal protein S23	0.59
P32970	CD70 antigen	0.59
Q01650	Large neutral amino acids transporter small subunit 1	0.59
P21266	Glutathione S-transferase Mu 3	0.59
Q14151	Scaffold attachment factor B2	0.59
P50914	60S ribosomal protein L14	0.59
Q16719	Kynureninase	0.60
Q9BVK6	Transmembrane emp24 domain-containing protein 9	0.60
P46782	40S ribosomal protein S5	0.60
Q9BS26	Endoplasmic reticulum resident protein 44	0.60
P38919	Eukaryotic initiation factor 4A-III	0.60
Q99714	3-hydroxyacyl-CoA dehydrogenase type-2	0.60
P35268	60S ribosomal protein L22	0.60
P46781	40S ribosomal protein S9	0.60
P31153	S-adenosylmethionine synthase isoform type-2	0.61
P51571	Translocon-associated protein subunit delta	0.61
P46778	60S ribosomal protein L21	0.62
P00403	Cytochrome c oxidase subunit 2	0.62
P62280	40S ribosomal protein S11	0.63
P52888	Thimet oligopeptidase	0.63
P06703	Protein S100-A6	0.63
P19338	Nucleolin	0.63
O00299	Chloride intracellular channel protein 1	0.63
P62249	40S ribosomal protein S16	0.63
P84243	Histone H3.3	0.63
Q9Y3U8	60S ribosomal protein L36	0.63
Q8NFJ5	Retinoic acid-induced protein 3	0.64
P00492	Hypoxanthine-guanine phosphoribosyltransferase	0.64
P61586	Transforming protein RhoA	0.64

<b>Q12905</b>	Interleukin enhancer-binding factor 2	0.64
<b>P26038</b>	Moesin	0.65
<b>Q00839</b>	Heterogeneous nuclear ribonucleoprotein U	0.65
<b>P07814</b>	Bifunctional glutamate/proline--tRNA ligase	0.66
<b>Q99650</b>	Oncostatin-M-specific receptor subunit beta	0.66
<b>P05198</b>	Eukaryotic translation initiation factor 2 subunit 1	0.66
<b>H7BZJ3</b>	Thioredoxin (Fragment)	0.66
<b>P49755</b>	Transmembrane emp24 domain-containing protein 10	0.66
<b>Q562R1</b>	Beta-actin-like protein 2	0.66
<b>P62263</b>	40S ribosomal protein S14	0.67
<b>Q07021</b>	Complement component 1 Q subcomponent-binding protein, mitochondrial	0.67
<b>I3L4N8</b>	Actin, cytoplasmic 2 (Fragment)	0.68
<b>P58107</b>	Epiplakin	0.68
<b>P13667</b>	Protein disulfide-isomerase A4	0.68
<b>P14625</b>	Endoplasmin	0.68
<b>P13639</b>	Elongation factor 2	0.69
<b>Q96AG4</b>	Leucine-rich repeat-containing protein 59	0.69
<b>P35579</b>	Myosin-9	0.71
<b>F5H867</b>	4F2 cell-surface antigen heavy chain (Fragment)	0.71
<b>Q9Y6M5</b>	Zinc transporter 1	0.71
<b>P11717</b>	Cation-independent mannose-6-phosphate receptor	0.72
<b>P21796</b>	Voltage-dependent anion-selective channel protein 1	0.72
<b>Q15365</b>	Poly(rC)-binding protein 1	0.72
<b>Q15738</b>	Sterol-4-alpha-carboxylate 3-dehydrogenase, decarboxylating	0.72
<b>Q969H8</b>	UPF0556 protein C19orf10	0.73
<b>P04406</b>	Glyceraldehyde-3-phosphate dehydrogenase	0.73
<b>P49327</b>	Fatty acid synthase	0.74
<b>Q9H3N1</b>	Thioredoxin-related transmembrane protein 1	0.74
<b>P27797</b>	Calreticulin	0.74
<b>P38646</b>	Stress-70 protein, mitochondrial	0.74
<b>P62805</b>	Histone H4	0.75

P04083	Annexin A1	0.75
Q9Y490	Talin-1	0.75
P46940	Ras GTPase-activating-like protein IQGAP1	0.75
P23284	Peptidyl-prolyl cis-trans isomerase B	0.76
Q15907	Ras-related protein Rab-11B	0.76
P05556	Integrin beta-1	0.77
Q06830	Peroxiredoxin-1	0.77
O95573	Long-chain-fatty-acid--CoA ligase 3	0.78
P00505	Aspartate aminotransferase, mitochondrial	0.78
P30043	Flavin reductase (NADPH)	0.78
P43007	Neutral amino acid transporter A	0.79
P55809	Succinyl-CoA:3-ketoacid coenzyme A transferase 1, mitochondrial	0.79
Q14764	Major vault protein	0.79
P54920	Alpha-soluble NSF attachment protein	0.81
Q13011	Delta(3,5)-Delta(2,4)-dienoyl-CoA isomerase, mitochondrial	0.81
P23458	Tyrosine-protein kinase JAK1	0.82
P12277	Creatine kinase B-type	0.82
Q9Y5M8	Signal recognition particle receptor subunit beta	0.82
P01891	HLA class I histocompatibility antigen, A-68 alpha chain	0.82
Q9UBS4	DnaJ homolog subfamily B member 11	0.83
O96008	Mitochondrial import receptor subunit TOM40 homolog	0.83
P07195	L-lactate dehydrogenase B chain	0.84
P84095	Rho-related GTP-binding protein RhoG	0.84
P68371	Tubulin beta-4B chain	0.86
O15173	Membrane-associated progesterone receptor component 2	0.86
Q96QK1	Vacuolar protein sorting-associated protein 35	0.87
Q01581	Hydroxymethylglutaryl-CoA synthase, cytoplasmic	0.87
B4DXP9	Alpha-centractin	0.88
P55072	Transitional endoplasmic reticulum ATPase	0.88
P61026	Ras-related protein Rab-10	0.91
P60842	Eukaryotic initiation factor 4A-I	0.92

<b>Q9UQ80</b>	Proliferation-associated protein 2G4	0.93
<b>Q9Y678</b>	Coatomer subunit gamma-1	0.93
<b>Q9H853</b>	Putative tubulin-like protein alpha-4B	0.94
<b>Q9Y696</b>	Chloride intracellular channel protein 4	0.95
<b>Q02790</b>	Peptidyl-prolyl cis-trans isomerase FKBP4	0.96
<b>Q8TEM1</b>	Nuclear pore membrane glycoprotein 210	0.96
<b>Q13162</b>	Peroxiredoxin-4	0.97
<b>Q9Y617</b>	Phosphoserine aminotransferase	1.00
<b>P56199</b>	Integrin alpha-1	1.00
<b>P04843</b>	Dolichyl-diphosphooligosaccharide--protein glycosyltransferase subunit 1	1.02
<b>P06733</b>	Alpha-enolase	1.02
<b>P07737</b>	Profilin-1	1.02
<b>P04792</b>	Heat shock protein beta-1	1.02
<b>P35579</b>	Myosin-9	1.03
<b>Q92616</b>	Translational activator GCN1	1.03
<b>O75396</b>	Vesicle-trafficking protein SEC22b	1.04
<b>Q04917</b>	14-3-3 protein eta	1.05
<b>Q8NF37</b>	Lysophosphatidylcholine acyltransferase 1	1.06
<b>Q9UBF2</b>	Coatomer subunit gamma-2	1.10
<b>P11169</b>	Solute carrier family 2, facilitated glucose transporter member 3	1.12
<b>Q9UL25</b>	Ras-related protein Rab-21	1.14
<b>P61981</b>	14-3-3 protein gamma	1.14
<b>Q15758</b>	Neutral amino acid transporter B(0)	1.16
<b>Q15181</b>	Inorganic pyrophosphatase	1.17
<b>P18669</b>	Phosphoglycerate mutase 1	1.18
<b>Q9UQE7</b>	Structural maintenance of chromosomes protein 3	1.28
<b>P11021</b>	78 kDa glucose-regulated protein	1.29
<b>P05141</b>	ADP/ATP translocase 2	1.33
<b>P05362</b>	Intercellular adhesion molecule 1	1.33
<b>P05161</b>	Ubiquitin-like protein ISG15	1.37
<b>P12236</b>	ADP/ATP translocase 3	1.38

<b>P41250</b>	Glycine--tRNA ligase	1.48
<b>P49588</b>	Alanine--tRNA ligase, cytoplasmic	1.55
<b>P35232</b>	Prohibitin	1.62
<b>O43175</b>	D-3-phosphoglycerate dehydrogenase	1.74
<b>P49411</b>	Elongation factor Tu, mitochondrial	1.82
<b>P42704</b>	Leucine-rich PPR motif-containing protein, mitochondrial	2.15
<b>P10809</b>	60 kDa heat shock protein, mitochondrial	2.73

\*: Heavy/Light means the protein expression ratio of PI4KII  $\alpha$  RNAi MCF-7 cells to Control cells.

**Table S2. Protein targets identified by SILAC and LC-MS/MS for control MCF-7 cells (sample 3) and combined inhibition MCF-7 cells (sample 4).**

Accession	Description	Heavy/Light*
<b>Q09666</b>	Neuroblast differentiation-associated protein AHNAK	0.21
<b>Q15459</b>	Splicing factor 3A subunit 1	0.28
<b>P16401</b>	Histone H1.5	0.29
<b>Q92945</b>	Far upstream element-binding protein 2	0.34
<b>Q9UNX4</b>	WD repeat-containing protein 3	0.34
<b>Q96PK6</b>	RNA-binding protein 14	0.36
<b>O94776</b>	Metastasis-associated protein MTA2	0.38
<b>Q05048</b>	Cleavage stimulation factor subunit 1	0.41
<b>O15213</b>	WD repeat-containing protein 46	0.43
<b>P14866</b>	Heterogeneous nuclear ribonucleoprotein L	0.43
<b>Q8NI36</b>	WD repeat-containing protein 36	0.44
<b>H7C267</b>	Pescadillo homolog (Fragment)	0.45
<b>Q15393</b>	Splicing factor 3B subunit 3	0.45
<b>P08670</b>	Vimentin	0.46
<b>P46063</b>	ATP-dependent DNA helicase Q1	0.46
<b>Q12996</b>	Cleavage stimulation factor subunit 3	0.46
<b>Q9BV38</b>	WD repeat-containing protein 18	0.46
<b>P15924</b>	Desmoplakin	0.47
<b>F6XZQ7</b>	Glutathione S-transferase Mu 2	0.48
<b>P26599</b>	Polypyrimidine tract-binding protein 1	0.48
<b>P11387</b>	DNA topoisomerase 1	0.48
<b>P09874</b>	Poly [ADP-ribose] polymerase 1	0.48
<b>Q01082</b>	Spectrin beta chain, non-erythrocytic 1	0.48
<b>P49327</b>	Fatty acid synthase	0.49
<b>P62750</b>	60S ribosomal protein L23a	0.50
<b>P08579</b>	U2 small nuclear ribonucleoprotein B	0.51
<b>P04792</b>	Heat shock protein beta-1	0.51

P12270	Nucleoprotein TPR	0.51
Q2TAY7	WD40 repeat-containing protein SMU1	0.51
Q15050	Ribosome biogenesis regulatory protein homolog	0.52
P12277	Creatine kinase B-type	0.53
P32119	Peroxiredoxin-2	0.54
P23246	Splicing factor, proline- and glutamine-rich	0.54
Q9BUQ8	Probable ATP-dependent RNA helicase DDX23	0.54
P17066	Heat shock 70 kDa protein 6	0.55
Q92499	ATP-dependent RNA helicase DDX1	0.55
Q9P258	Protein RCC2	0.56
Q9Y224	UPF0568 protein C14orf166	0.56
P13010	X-ray repair cross-complementing protein 5	0.56
P49736	DNA replication licensing factor MCM2	0.56
P80723	Brain acid soluble protein 1	0.56
P38919	Eukaryotic initiation factor 4A-III	0.57
Q12905	Interleukin enhancer-binding factor 2	0.57
Q9NS69	Mitochondrial import receptor subunit TOM22 homolog	0.57
Q15269	Periodic tryptophan protein 2 homolog	0.58
O75533	Splicing factor 3B subunit 1	0.58
Q13907-2	Isoform 2 of Isopentenyl-diphosphate Delta-isomerase 1	0.59
Q08945	FACT complex subunit SSRP1	0.59
Q9GZL7	Ribosome biogenesis protein WDR12	0.59
O60264	SWI/SNF-related matrix-associated actin-dependent regulator of chromatin subfamily A member 5	0.602
P62304	Small nuclear ribonucleoprotein E	0.60
Q15397	Pumilio domain-containing protein KIAA0020	0.61
Q8IY81	pre-rRNA processing protein FTSJ3	0.61
O43809	Cleavage and polyadenylation specificity factor subunit 5	0.61
P52907	F-actin-capping protein subunit alpha-1	0.61
Q00839	Heterogeneous nuclear ribonucleoprotein U	0.61
Q08211	ATP-dependent RNA helicase A	0.62

<b>Q7KZF4</b>	Staphylococcal nuclease domain-containing protein 1	0.62
<b>Q9Y3T9</b>	Nucleolar complex protein 2 homolog	0.63
<b>Q14683</b>	Structural maintenance of chromosomes protein 1A	0.63
<b>Q05639</b>	Elongation factor 1-alpha 2	0.64
<b>O75691</b>	Small subunit processome component 20 homolog	0.64
<b>Q8IWA0</b>	WD repeat-containing protein 75	0.64
<b>Q9Y678</b>	Coatomer subunit gamma-1	0.64
<b>Q13151</b>	Heterogeneous nuclear ribonucleoprotein A0	0.65
<b>O15144</b>	Actin-related protein 2/3 complex subunit 2	0.65
<b>P15121</b>	Aldose reductase	0.65
<b>Q96GQ7</b>	Probable ATP-dependent RNA helicase DDX27	0.65
<b>P52292</b>	Importin subunit alpha-2	0.66
<b>P56182</b>	Ribosomal RNA processing protein 1 homolog A	0.66
<b>Q12788</b>	Transducin beta-like protein 3	0.66
<b>P23284</b>	Peptidyl-prolyl cis-trans isomerase B	0.66
<b>O75531</b>	Barrier-to-autointegration factor	0.67
<b>Q9Y2P8</b>	RNA 3'-terminal phosphate cyclase-like protein	0.67
<b>Q9NVP1</b>	ATP-dependent RNA helicase DDX18	0.67
<b>P16083</b>	Ribosyldihydronicotinamide dehydrogenase [quinone]	0.68
<b>P20700</b>	Lamin-B1	0.68
<b>Q9Y5B9</b>	FACT complex subunit SPT16	0.68
<b>Q06203</b>	Amidophosphoribosyltransferase	0.69
<b>O00148</b>	ATP-dependent RNA helicase DDX39A	0.69
<b>O43818</b>	U3 small nucleolar RNA-interacting protein 2	0.69
<b>Q03701</b>	CCAAT/enhancer-binding protein zeta	0.70
<b>Q6P2Q9</b>	Pre-mRNA-processing-splicing factor 8	0.70
<b>P37802</b>	Transgelin-2	0.71
<b>P30740</b>	Leukocyte elastase inhibitor	0.71
<b>P08574</b>	Cytochrome c1, heme protein, mitochondrial	0.72
<b>Q9UQE7</b>	Structural maintenance of chromosomes protein 3	0.72
<b>Q9Y2W1</b>	Thyroid hormone receptor-associated protein 3	0.72

<b>Q14690</b>	Protein RRP5 homolog	0.72
<b>P53004</b>	Biliverdin reductase A	0.72
<b>Q15365</b>	Poly(rC)-binding protein 1	0.72
<b>Q9Y5J1</b>	U3 small nucleolar RNA-associated protein 18 homolog	0.73
<b>P35237</b>	Serpin B6	0.74
<b>P36871</b>	Phosphoglucomutase-1	0.74
<b>Q8TDN6</b>	Ribosome biogenesis protein BRX1 homolog	0.74
<b>O00232</b>	26S proteasome non-ATPase regulatory subunit 12	0.74
<b>P08238</b>	Heat shock protein HSP 90-beta	0.75
<b>E9PBF6</b>	Lamin-B1	0.75
<b>Q15738</b>	Sterol-4-alpha-carboxylate 3-dehydrogenase, decarboxylating	0.75
<b>O60812</b>	Heterogeneous nuclear ribonucleoprotein C-like 1	0.75
<b>P05783</b>	Keratin, type I cytoskeletal 18	0.76
<b>O00567</b>	Nucleolar protein 56	0.77
<b>P04080</b>	Cystatin-B	0.77
<b>Q9Y295</b>	Developmentally-regulated GTP-binding protein 1	0.78
<b>Q92979</b>	Ribosomal RNA small subunit methyltransferase NEP1	0.78
<b>P31949</b>	Protein S100-A11	0.78
<b>Q96DG6</b>	Carboxymethylenebutenolidase homolog	0.78
<b>Q9NV31</b>	U3 small nucleolar ribonucleoprotein protein IMP3	0.78
<b>P05386</b>	60S acidic ribosomal protein P1	0.78
<b>P09382</b>	Galectin-1	0.78
<b>Q86U38</b>	Nucleolar protein 9	0.78
<b>P25815</b>	Protein S100-P	0.79
<b>Q02790</b>	Peptidyl-prolyl cis-trans isomerase FKBP4	0.79
<b>Q9BYZ2</b>	L-lactate dehydrogenase A-like 6B	0.79
<b>P49207</b>	60S ribosomal protein L34	0.79
<b>Q9NYH9</b>	U3 small nucleolar RNA-associated protein 6 homolog	0.79
<b>P22314</b>	Ubiquitin-like modifier-activating enzyme 1	0.81
<b>P26038</b>	Moesin	0.81
<b>P26447</b>	Protein S100-A4	0.81

<b>O43143</b>	Putative pre-mRNA-splicing factor ATP-dependent RNA helicase DHX15	0.81
<b>P48047</b>	ATP synthase subunit O, mitochondrial	0.82
<b>P29317</b>	Ephrin type-A receptor 2	0.82
<b>P31947</b>	14-3-3 protein sigma	0.82
<b>P30086</b>	Phosphatidylethanolamine-binding protein 1	0.82
<b>P34932</b>	Heat shock 70 kDa protein 4	0.83
<b>P58107</b>	Epiplakin	0.83
<b>Q9BZK3</b>	Putative nascent polypeptide-associated complex subunit alpha-like protein	0.83
<b>Q9Y2X3</b>	Nucleolar protein 58	0.83
<b>O60287</b>	Nucleolar pre-ribosomal-associated protein 1	0.83
<b>F5H0T1</b>	Stress-induced-phosphoprotein 1	0.83
<b>P28066</b>	Proteasome subunit alpha type-5	0.84
<b>P28072</b>	Proteasome subunit beta type-6	0.84
<b>P35232</b>	Prohibitin	0.84
<b>P62424</b>	60S ribosomal protein L7a	0.84
<b>P49721</b>	Proteasome subunit beta type-2	0.85
<b>P62854</b>	40S ribosomal protein S26	0.86
<b>P18669</b>	Phosphoglycerate mutase 1	0.86
<b>P00491</b>	Purine nucleoside phosphorylase	0.86
<b>P26373</b>	60S ribosomal protein L13	0.86
<b>P07195</b>	L-lactate dehydrogenase B chain	0.87
<b>P21796</b>	Voltage-dependent anion-selective channel protein 1	0.87
<b>Q01581</b>	Hydroxymethylglutaryl-CoA synthase, cytoplasmic	0.87
<b>Q92621</b>	Nuclear pore complex protein Nup205	0.88
<b>P62081</b>	40S ribosomal protein S7	0.88
<b>Q06830</b>	Peroxiredoxin-1	0.88
<b>E9PCK7</b>	RRP12-like protein	0.89
<b>Q99436</b>	Proteasome subunit beta type-7	0.89
<b>Q99497</b>	Protein DJ-1	0.89
<b>P31153</b>	S-adenosylmethionine synthase isoform type-2	0.89
<b>P06744</b>	Glucose-6-phosphate isomerase	0.90

<b>Q9UL46</b>	Proteasome activator complex subunit 2	0.90
<b>P20618</b>	Proteasome subunit beta type-1	0.90
<b>P30041</b>	Peroxiredoxin-6	0.90
<b>P30050</b>	60S ribosomal protein L12	0.90
<b>P78527</b>	DNA-dependent protein kinase catalytic subunit	0.90
<b>Q9NV06</b>	DDB1- and CUL4-associated factor 13	0.90
<b>Q9Y230</b>	RuvB-like 2	0.90
<b>P06703</b>	Protein S100-A6	0.91
<b>P25705</b>	ATP synthase subunit alpha, mitochondrial	0.91
<b>P49257</b>	Protein ERGIC-53	0.91
<b>A6NDG6</b>	Phosphoglycolate phosphatase	0.91
<b>P61586</b>	Transforming protein RhoA	0.91
<b>P62263</b>	40S ribosomal protein S14	0.91
<b>Q562R1</b>	Beta-actin-like protein 2	0.91
<b>P23921</b>	Ribonucleoside-diphosphate reductase large subunit	0.92
<b>P51571</b>	Translocon-associated protein subunit delta	0.92
<b>P04406</b>	Glyceraldehyde-3-phosphate dehydrogenase	0.92
<b>P42166</b>	Lamina-associated polypeptide 2, isoform alpha	0.92
<b>P62277</b>	40S ribosomal protein S13	0.92
<b>P06733</b>	Alpha-enolase	0.92
<b>P42766</b>	60S ribosomal protein L35	0.92
<b>O00299</b>	Chloride intracellular channel protein 1	0.92
<b>O95336</b>	6-phosphogluconolactonase	0.92
<b>P07737</b>	Profilin-1	0.92
<b>P13639</b>	Elongation factor 2	0.92
<b>O75131</b>	Copine-3	0.93
<b>P08754</b>	Guanine nucleotide-binding protein G(k) subunit alpha	0.93
<b>P47914</b>	60S ribosomal protein L29	0.93
<b>P08708</b>	40S ribosomal protein S17	0.93
<b>P25787</b>	Proteasome subunit alpha type-2	0.93
<b>P30084</b>	Enoyl-CoA hydratase, mitochondrial	0.93

<b>P0CG39</b>	POTE ankyrin domain family member J	0.94
<b>P35268</b>	60S ribosomal protein L22	0.94
<b>P41091</b>	Eukaryotic translation initiation factor 2 subunit 3	0.94
<b>Q01813</b>	6-phosphofructokinase type C	0.94
<b>Q8N726</b>	Cyclin-dependent kinase inhibitor 2A, isoform 4	0.94
<b>P18124</b>	60S ribosomal protein L7	0.94
<b>P25398</b>	40S ribosomal protein S12	0.94
<b>Q9H9B4</b>	Sideroflexin-1	0.94
<b>P46940</b>	Ras GTPase-activating-like protein IQGAP1	0.95
<b>Q9BQ39</b>	ATP-dependent RNA helicase DDX50	0.95
<b>P24534</b>	Elongation factor 1-beta	0.95
<b>P12004</b>	Proliferating cell nuclear antigen	0.96
<b>Q16719</b>	Kynureninase	0.96
<b>P52597</b>	Heterogeneous nuclear ribonucleoprotein F	0.96
<b>P62269</b>	40S ribosomal protein S18	0.96
<b>P61254</b>	60S ribosomal protein L26	0.96
<b>Q14019</b>	Coactosin-like protein	0.96
<b>F5H1L6</b>	Serine/threonine-protein phosphatase PP1-alpha catalytic subunit (Fragment)	0.97
<b>P16152</b>	Carbonyl reductase [NADPH] 1	0.97
<b>P46777</b>	60S ribosomal protein L5	0.97
<b>Q15181</b>	Inorganic pyrophosphatase	0.97
<b>P61313</b>	60S ribosomal protein L15	0.98
<b>Q9HAV0</b>	Guanine nucleotide-binding protein subunit beta-4	0.99
<b>P51153</b>	Ras-related protein Rab-13	0.99
<b>O75489</b>	NADH dehydrogenase [ubiquinone] iron-sulfur protein 3, mitochondrial	0.99
<b>I3L4N8</b>	Actin, cytoplasmic 2 (Fragment)	0.99
<b>P00367</b>	Glutamate dehydrogenase 1, mitochondrial	0.99
<b>Q9Y3U8</b>	60S ribosomal protein L36	0.99
<b>P62249</b>	40S ribosomal protein S16	0.99
<b>E7ES52</b>	Cyclin-dependent kinase inhibitor 1B	1.00
<b>P21266</b>	Glutathione S-transferase Mu 3	1.00

P60903	Protein S100-A10	1.00
Q96AG4	Leucine-rich repeat-containing protein 59	1.00
Q8TD16-2	Isoform 2 of Protein bicaudal D homolog 2	1.00
P00441	Superoxide dismutase [Cu-Zn]	1.01
P62280	40S ribosomal protein S11	1.01
P62266	40S ribosomal protein S23	1.01
Q9UBI6	Guanine nucleotide-binding protein G(I)/G(S)/G(O) subunit gamma-12	1.01
P46782	40S ribosomal protein S5	1.02
P55072	Transitional endoplasmic reticulum ATPase	1.02
P15311	Ezrin	1.02
P53985	Monocarboxylate transporter 1	1.02
P53999	Activated RNA polymerase II transcriptional coactivator p15	1.02
P62701	40S ribosomal protein S4, X isoform	1.02
P23458	Tyrosine-protein kinase JAK1	1.03
Q92616	Translational activator GCN1	1.03
P61353	60S ribosomal protein L27	1.03
P62906	60S ribosomal protein L10a	1.03
P31689	DnaJ homolog subfamily A member 1	1.03
P27348	14-3-3 protein theta	1.04
P30041	Peroxiredoxin-6	1.04
P23526	Adenosylhomocysteinase	1.04
P50914	60S ribosomal protein L14	1.04
P14625	Endoplasmin	1.04
P62841	40S ribosomal protein S15	1.04
Q9NR45	Sialic acid synthase	1.04
Q969H8	UPF0556 protein C19orf10	1.05
P28070	Proteasome subunit beta type-4	1.05
P43121	Cell surface glycoprotein MUC18	1.05
Q969Q0	60S ribosomal protein L36a-like	1.05
P35579	Myosin-9	1.05
O76003	Glutaredoxin-3	1.06

P55795	Heterogeneous nuclear ribonucleoprotein H2	1.06
P61006	Ras-related protein Rab-8A	1.07
O00487	26S proteasome non-ATPase regulatory subunit 14	1.07
O43707	Alpha-actinin-4	1.07
Q99714	3-hydroxyacyl-CoA dehydrogenase type-2	1.08
O75643	U5 small nuclear ribonucleoprotein 200 kDa helicase	1.08
P46778	60S ribosomal protein L21	1.08
Q04917	14-3-3 protein eta	1.08
P40429	60S ribosomal protein L13a	1.09
Q12907	Vesicular integral-membrane protein VIP36	1.09
Q15006	ER membrane protein complex subunit 2	1.09
P68371	Tubulin beta-4B chain	1.10
Q9H3N1	Thioredoxin-related transmembrane protein 1	1.10
H0YKF0	Electron transfer flavoprotein subunit alpha, mitochondrial (Fragment)	1.11
P37837	Transaldolase	1.11
P07814	Bifunctional glutamate/proline--tRNA ligase	1.11
P05198	Eukaryotic translation initiation factor 2 subunit 1	1.11
Q01105	Protein SET	1.12
P62805	Histone H4	1.12
Q9UMS4	Pre-mRNA-processing factor 19	1.12
P05556	Integrin beta-1	1.12
P22087	rRNA 2'-O-methyltransferase fibrillarin	1.13
Q16795	NADH dehydrogenase [ubiquinone] 1 alpha subcomplex subunit 9, mitochondrial	1.14
P19338	Nucleolin	1.14
Q9H4A6	Golgi phosphoprotein 3	1.14
P04083	Annexin A1	1.15
Q9Y5M8	Signal recognition particle receptor subunit beta	1.15
P55809	Succinyl-CoA:3-ketoacid coenzyme A transferase 1, mitochondrial	1.15
P84243	Histone H3.3	1.15
Q13641	Trophoblast glycoprotein	1.16
P11166	Solute carrier family 2, facilitated glucose transporter member 1	1.16

P02792	Ferritin light chain	1.16
Q14764	Major vault protein	1.17
P48637	Glutathione synthetase	1.17
P30040	Endoplasmic reticulum resident protein 29	1.188
Q10589	Bone marrow stromal antigen 2	1.19
O96008	Mitochondrial import receptor subunit TOM40 homolog	1.19
P31930	Cytochrome b-c1 complex subunit 1, mitochondrial	1.19
O75874	Isocitrate dehydrogenase [NADP] cytoplasmic	1.19
P54920	Alpha-soluble NSF attachment protein	1.19
Q71DI3	Histone H3.2	1.20
P00505	Aspartate aminotransferase, mitochondrial	1.20
P60842	Eukaryotic initiation factor 4A-I	1.21
P50148	Guanine nucleotide-binding protein G(q) subunit alpha	1.22
P00966	Argininosuccinate synthase	1.22
P04843	Dolichyl-diphosphooligosaccharide--protein glycosyltransferase subunit 1	1.23
Q96QK1	Vacuolar protein sorting-associated protein 35	1.24
O14980	Exportin-1	1.24
Q9BVK6	Transmembrane emp24 domain-containing protein 9	1.24
Q13011	Delta(3,5)-Delta(2,4)-dienoyl-CoA isomerase, mitochondrial	1.25
P27797	Calreticulin	1.26
Q01650	Large neutral amino acids transporter small subunit 1	1.26
H7BZJ3	Thioredoxin (Fragment)	1.28
P12236	ADP/ATP translocase 3	1.28
P00492	Hypoxanthine-guanine phosphoribosyltransferase	1.29
P30043	Flavin reductase (NADPH)	1.29
O95373	Importin-7	1.31
O60568	Procollagen-lysine,2-oxoglutarate 5-dioxygenase 3	1.32
P62745	Rho-related GTP-binding protein RhoB	1.34
Q07065	Cytoskeleton-associated protein 4	1.34
P10599	Thioredoxin	1.35
P40261	Nicotinamide N-methyltransferase	1.35

<b>Q16740</b>	Putative ATP-dependent Clp protease proteolytic subunit, mitochondrial	1.36
<b>O95573</b>	Long-chain-fatty-acid--CoA ligase 3	1.37
<b>Q96HE7</b>	ERO1-like protein alpha	1.37
<b>P48556</b>	26S proteasome non-ATPase regulatory subunit 8	1.38
<b>P01891</b>	HLA class I histocompatibility antigen, A-68 alpha chain	1.38
<b>P61981</b>	14-3-3 protein gamma	1.39
<b>P67936</b>	Tropomyosin alpha-4 chain	1.40
<b>P61106</b>	Ras-related protein Rab-14	1.41
<b>O43681</b>	ATPase ASNA1	1.42
<b>O75396</b>	Vesicle-trafficking protein SEC22b	1.42
<b>H7C3T4</b>	Peroxiredoxin-4 (Fragment)	1.44
<b>P05141</b>	ADP/ATP translocase 2	1.44
<b>Q04837</b>	Single-stranded DNA-binding protein, mitochondrial	1.47
<b>Q9UHV9</b>	Prefoldin subunit 2	1.53
<b>P13667</b>	Protein disulfide-isomerase A4	1.55
<b>Q6UB35</b>	Monofunctional C1-tetrahydrofolate synthase, mitochondrial	1.60
<b>Q8NFH3</b>	Nucleoporin Nup43	1.60
<b>Q15645</b>	Pachytene checkpoint protein 2 homolog	1.62
<b>Q8IV08</b>	Phospholipase D3	1.62
<b>P24752</b>	Acetyl-CoA acetyltransferase, mitochondrial	1.64
<b>P05161</b>	Ubiquitin-like protein ISG15	1.66
<b>Q14566</b>	DNA replication licensing factor MCM6	1.71
<b>P05362</b>	Intercellular adhesion molecule 1	1.72
<b>O15173</b>	Membrane-associated progesterone receptor component 2	1.78
<b>Q8TEM1</b>	Nuclear pore membrane glycoprotein 210	1.81
<b>P21266</b>	Glutathione S-transferase Mu 3	1.84
<b>P61026</b>	Ras-related protein Rab-10	1.87
<b>P49755</b>	Transmembrane emp24 domain-containing protein 10	1.88
<b>P11021</b>	78 kDa glucose-regulated protein	1.97
<b>P49411</b>	Elongation factor Tu, mitochondrial	2.00
<b>P41250</b>	Glycine--tRNA ligase	2.17

<b>P42704</b>	Leucine-rich PPR motif-containing protein, mitochondrial	2.27
<b>Q15758</b>	Neutral amino acid transporter B(0)	2.32
<b>P24539</b>	ATP synthase subunit b, mitochondrial	2.48
<b>P40939</b>	Trifunctional enzyme subunit alpha, mitochondrial	2.50
<b>Q07021</b>	Complement component 1 Q subcomponent-binding protein, mitochondrial	2.52
<b>O43175</b>	D-3-phosphoglycerate dehydrogenase	2.52=3
<b>P10809</b>	60 kDa heat shock protein, mitochondrial	<b>2.67</b>
<b>Q9Y617</b>	Phosphoserine aminotransferase	2.87]
<b>P49588</b>	Alanine--tRNA ligase, cytoplasmic	2.88]
<b>Q14204</b>	Cytoplasmic dynein 1 heavy chain 1	3.04]
<b>P38646</b>	Stress-70 protein, mitochondrial	3.48]
<b>E9PPU0</b>	Epiplakin	4.29

\*: Heavy/Light means the protein expression ratio of PI4KII  $\alpha$  and EGFR dual inhibited MCF-7 cells to Control cells.