

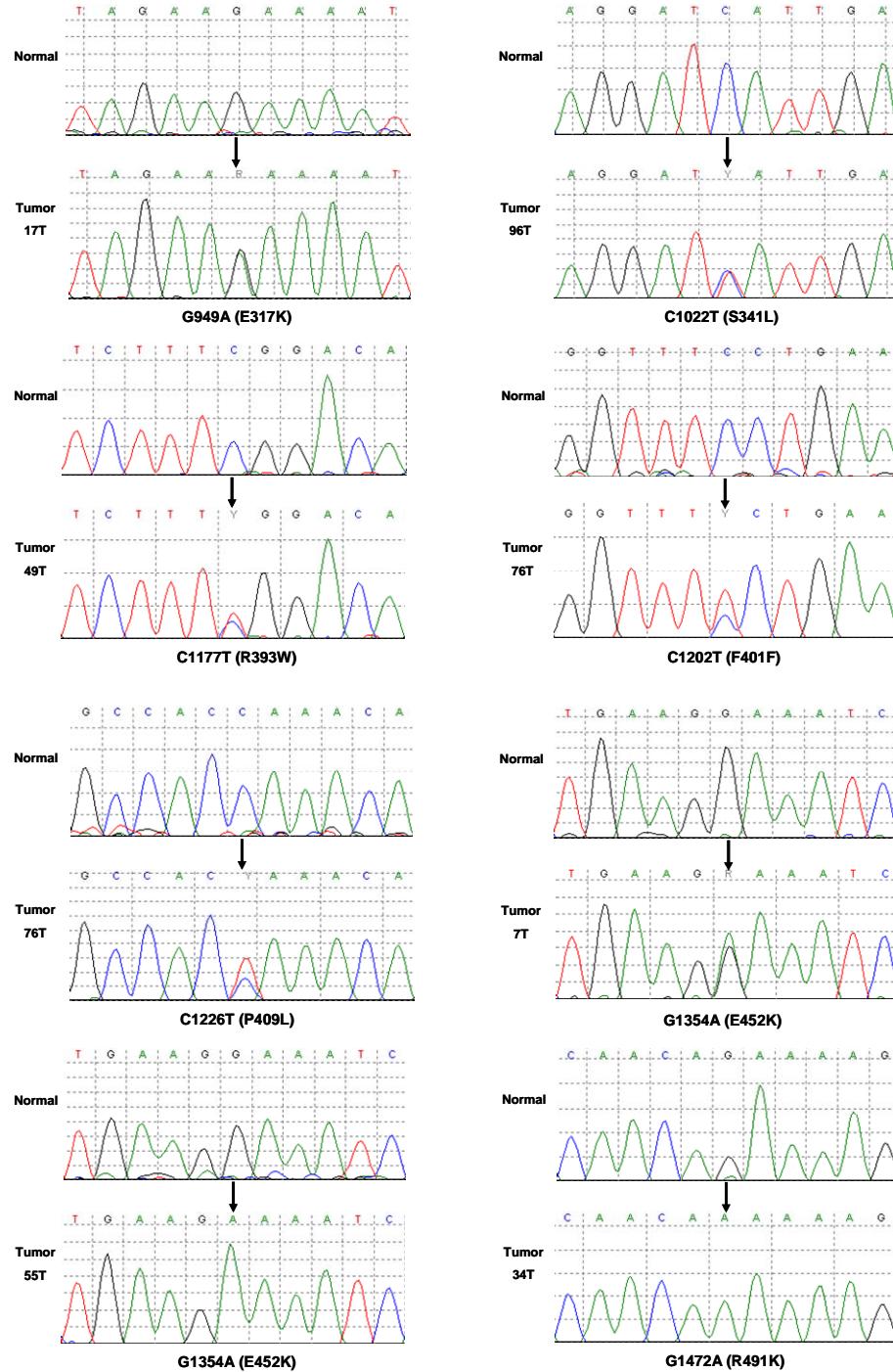
Supplementary Material:

Analysis of the Tyrosine Kinome in Melanoma Reveals Recurrent Mutations in *ERBB4*

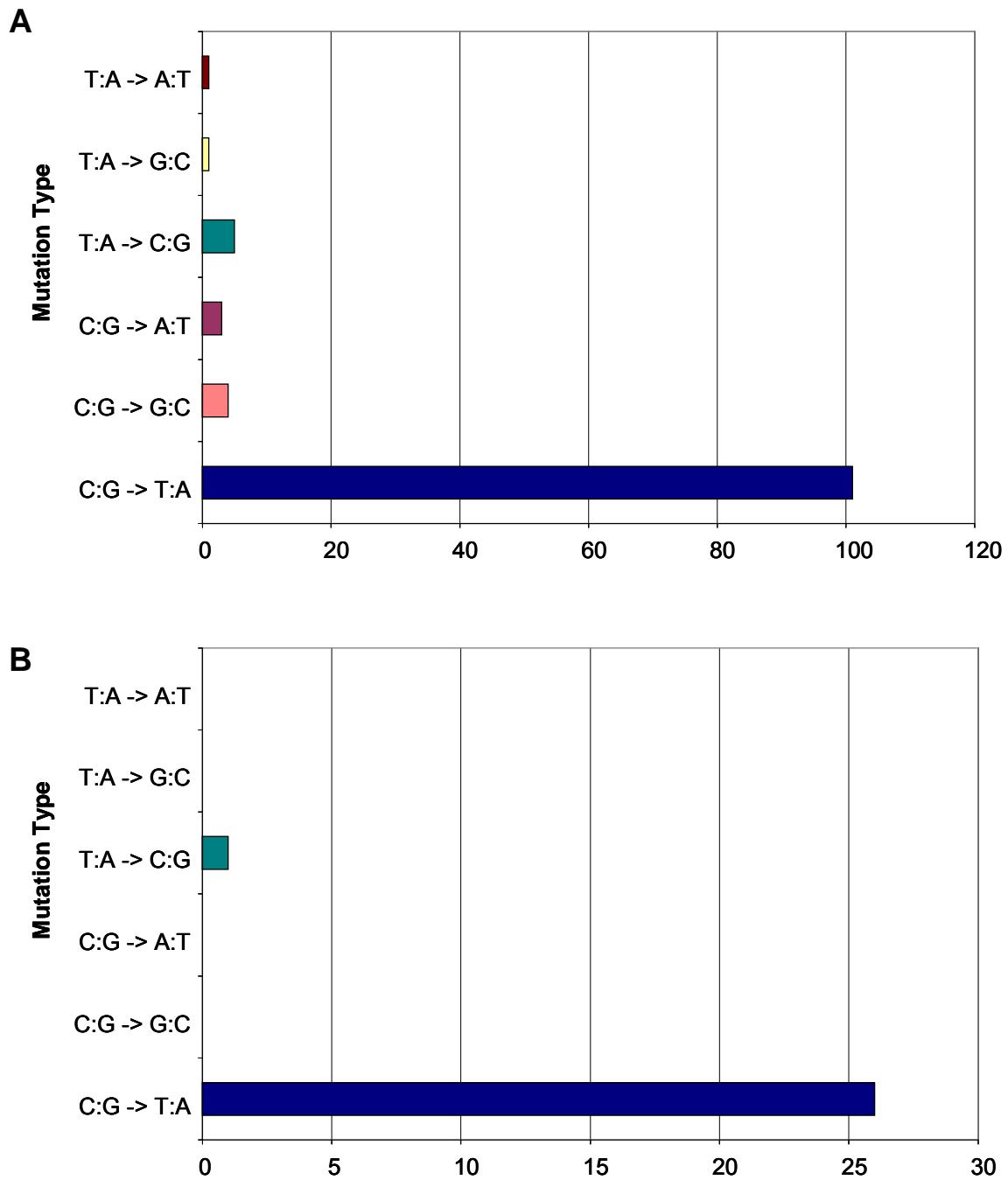
Todd D. Prickett, Neena S. Agrawal, Xiaomu Wei, Kristin E. Yates, Jimmy C. Lin, John Wunderlich, Julia C. Cronin, Pedro Cruz, NISC Comparative Sequencing Program, Steven A. Rosenberg, Yardena Samuels*

* Corresponding author: Yardena Samuels, National Human Genome Research Institute, NIH, 50 South Drive, MSC 8000, Building 50, Room 5140, Bethesda MD 20892-8000, Phone: 301-451-2628, Fax: 301-480-9864, Email: samuelsy@mail.nih.gov

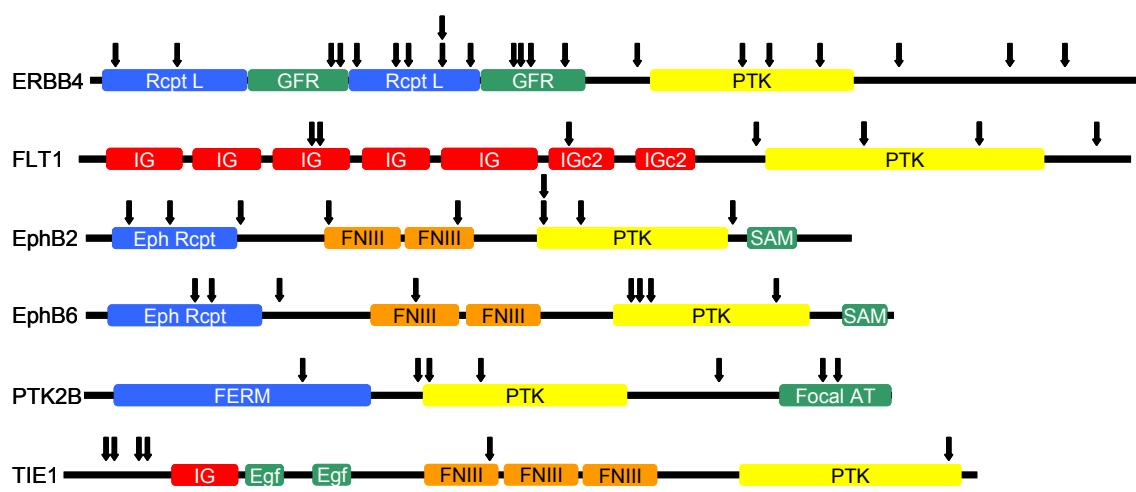
Supplementary Figures



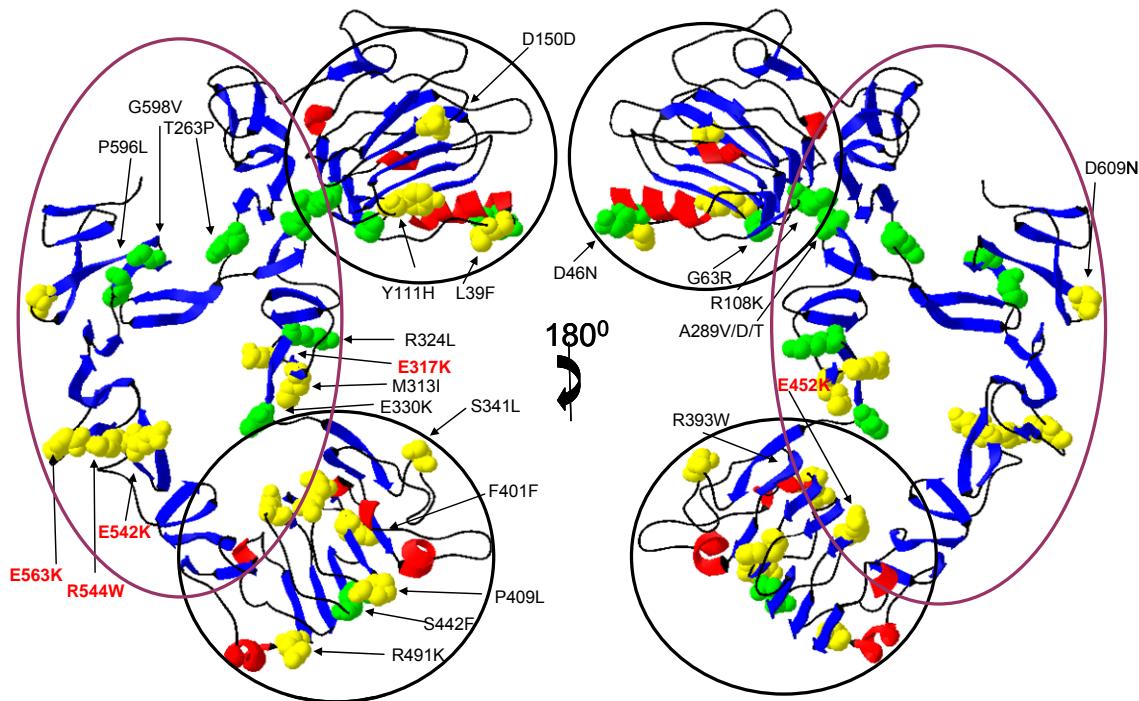
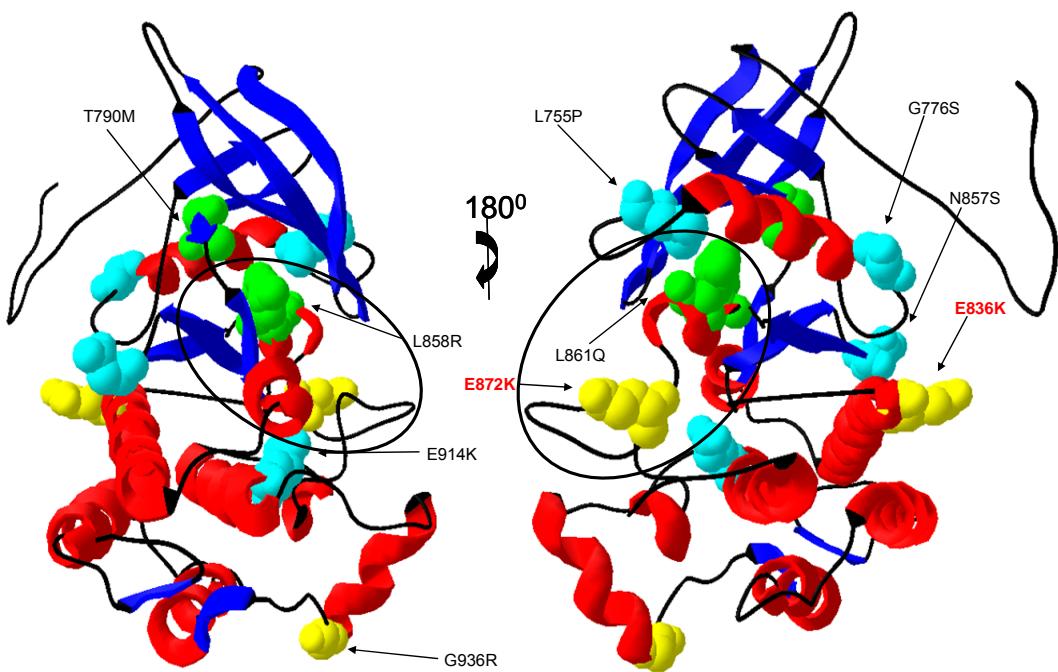
Supplementary Figure 1. Detection of mutations in ERBB4. Representative examples of mutations in ERBB4. In each case, the top sequence chromatogram was obtained from normal tissue and the lower sequence chromatogram from the indicated tumors. Arrows indicate the location of missense mutations. The nucleotide and amino acid alterations are indicated below the chromatograms.



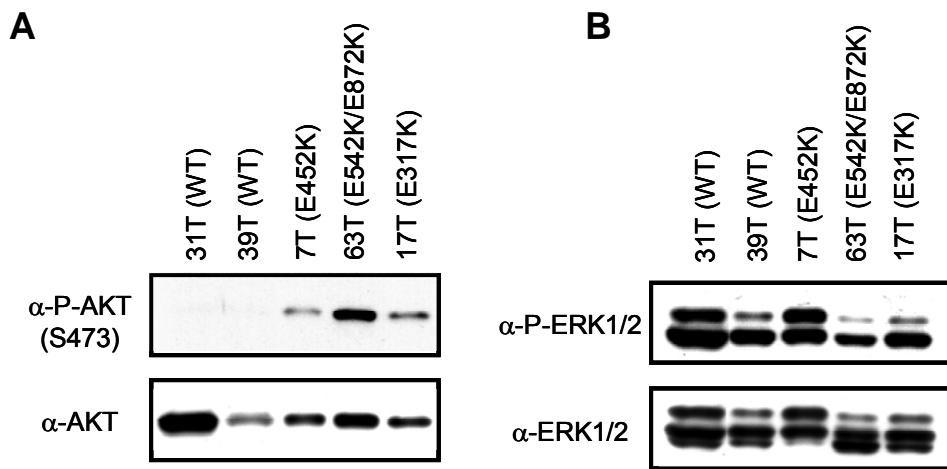
Supplementary Figure 2. Mutation spectra of single base pair substitutions.
A. Kinome mutation spectrum. The number of each of the six classes of base substitutions resulting in nonsynonymous changes in the kinome screen is shown. B. Mutation spectra of single base pair substitutions in ERBB4. The number of each of the six classes of base substitutions resulting in nonsynonymous changes in ERBB4 is shown.



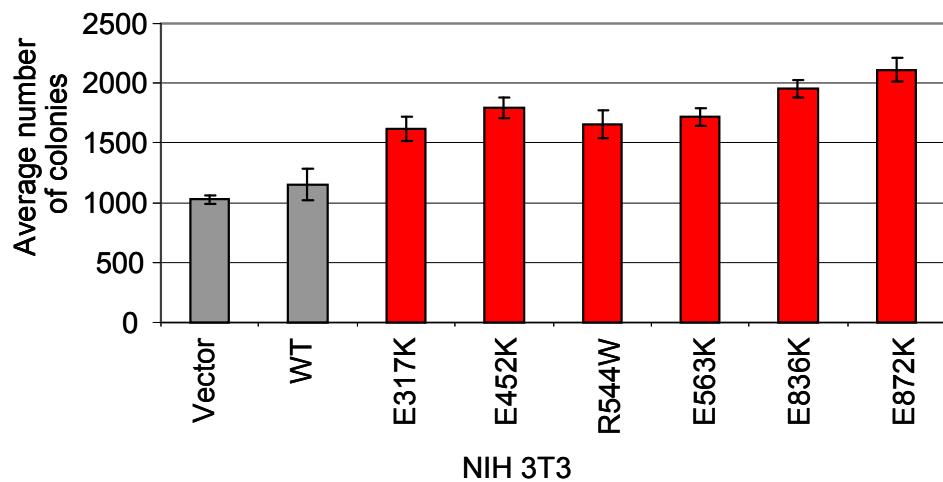
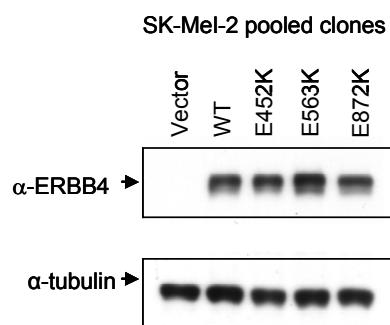
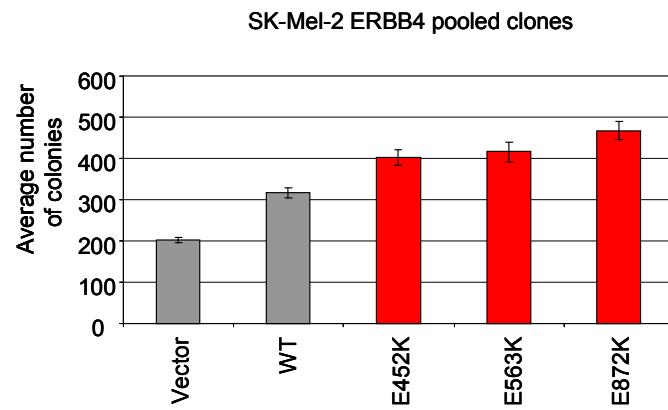
Supplementary Figure 3. Distribution of mutations in ERBB4, FLT1, EphB2, EphB6, PTK2B, and TIE1. Black arrows indicate positions of nonsynonymous mutations and boxes represent functional domains (Rcpt L, Receptor L; GFR, Growth Factor Receptor; PTK, Protein Tyrosine Kinase; IG, Immunoglobulin; IGc2, Immunoglobulin C-2 Type; Eph Rcpt, Ephrin Receptor; FNIII, Fibronectin Type III; SAM, Sterile Alpha Motif; FERM, Protein 4.1, Ezrin, Radixin, Moesin (FERM) Domain; Focal AT, Focal Adhesion Targeting Region)

A**B**

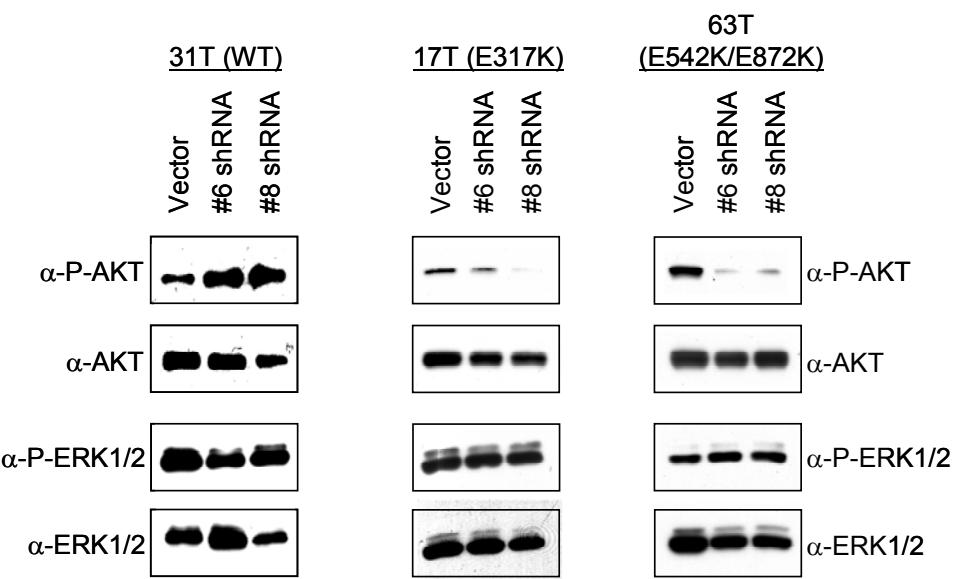
Supplementary Figure 4. Ribbon diagrams of ERBB4. A. Distribution of ERBB4 mutations in a ribbon diagram of its extracellular domain (PDB ID 2ahx). The two views are related to one another by a 180° rotation about a vertical axis. α -helices are shown in red, β -sheets are shown in blue, and loops are shown in black. All mutations are represented as space-filling molecules. ERBB4 mutations are shown in yellow, and the equivalent positions of known activating EGFR mutations are shown in green. The receptor L domain is denoted by black circles, the growth factor receptor domain is denoted by purple circles and cloned ERBB4 mutants are highlighted in red. B. Distribution of ERBB4 mutations in a ribbon of its kinase domain (PDB ID 2r4b). The two views are related to one another by a 180° rotation about a vertical axis. α -helices are shown in red, β -sheets are shown in blue, and loops are shown in black. All mutations are represented as space-filling molecules. ERBB4 mutations are shown in yellow, equivalent positions of known ERBB2 mutations are shown in cyan, and the equivalent positions of known activating EGFR mutations are shown in green. The activation loop is denoted by black circles. In A and B, Images were created by SWISS-MODEL. Determination of the location of previously reported EGFR and ERBB2 mutations is described in the experimental procedures and cloned ERBB4 mutants are highlighted in red.



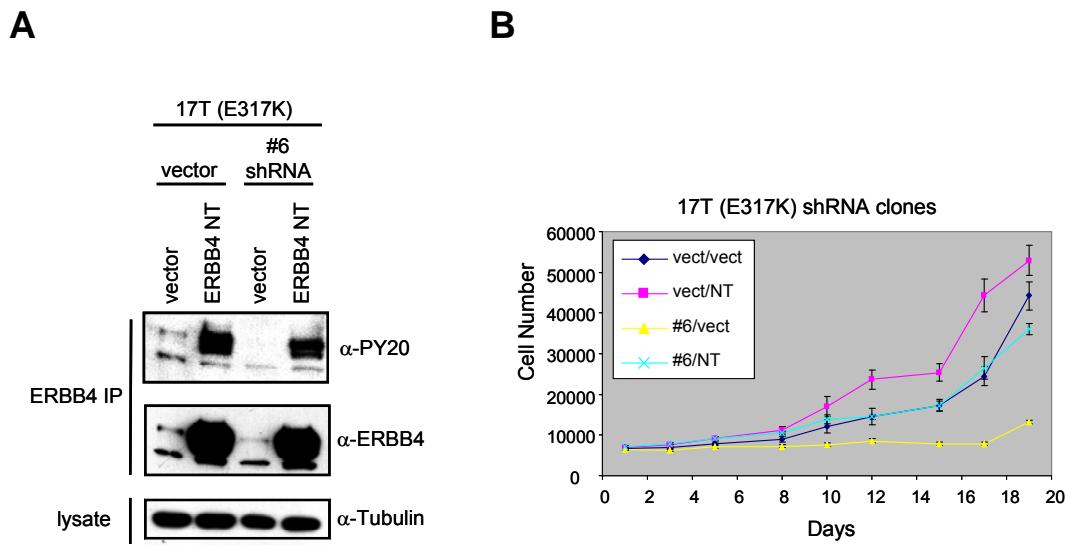
Supplementary Figure 5. Effects of ERBB4 mutation on AKT and ERK phosphorylation. Melanoma cell lines, containing either WT or mutant ERBB4, were harvested and analyzed by immunoblot. Shown are immunoblots of lysates probed with the indicated antibodies (α -P-ERK1/2 (recognizes phosphorylation of T202 and Y204 on ERK1 and T185 and Y187 on ERK2).

A**B****C**

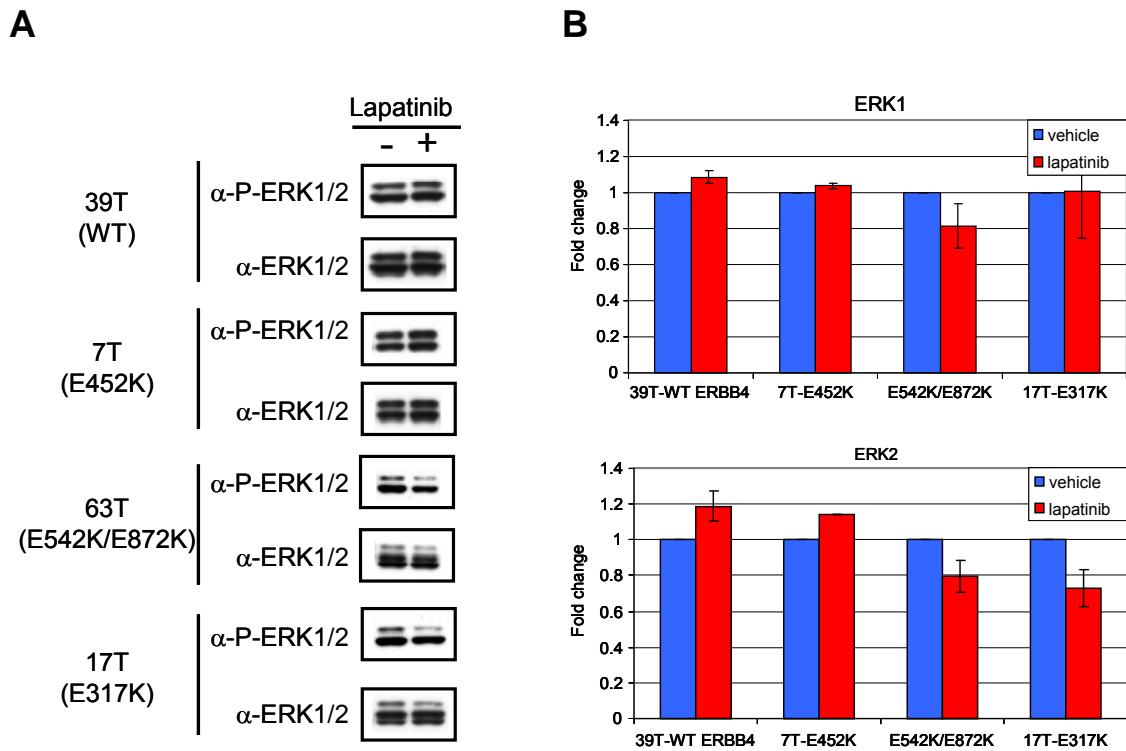
Supplementary Figure 6. Effect of ERBB4 mutations on cell growth in NIH 3T3 and SK-Mel-2 cells. A. Growth of NIH 3T3 cells expressing vector, WT ERBB4 or various ERBB4 missense mutants in soft agar. The graph indicates the number of colonies after 14 days. B. Detection of ERBB4 protein expression in stable transfectants of SK-Mel-2 melanoma cells by immunoblot analysis, lysates from the different clones stably transfected with an empty vector, human WT ERBB4 or the indicated ERBB4 mutants were immunoprecipitated and immunoblotted with ERBB4 antibody. C. Anchorage-independent proliferation of SK-Mel-2 cell clones expressing the indicated constructs was assessed by measuring colony formation in soft agar. Graph indicates number of colonies observed after 14 days of growth.



Supplementary Figure 7. Knockdown of ERBB4 protein causes reduced activation of the AKT pathway but not of the ERK pathway. Melanoma cell lines, containing either WT or mutant ERBB4, were harvested and analyzed by western blot. Shown are immunoblots of lysates probed with the indicated antibodies (α -P-ERK1/2 (recognizes phosphorylation of T202 and Y204 on ERK1 and T185 and Y187 on ERK2).



Supplementary Figure 8. Rescue of oncogene dependence by exogenous non-targetable ERBB4. A. Melanoma cells harboring mutant ERBB4 and stably expressing control or ERBB4 shRNA #6 transduced with either vector or non targetable (NT) ERBB4 were analyzed by immunoprecipitation of lysates with α -ERBB4. Samples were analyzed by immunoblotting with the indicated antibodies. As a loading control lysates were immunoblotted with α -Tubulin. B. Melanoma cells expressing vector or the ERBB4 shRNA #6 transduced with a vector or NT ERBB4 were evaluated for cell proliferation by measuring the average cell number at each time point by determining DNA content using SYBR Green I.



Supplementary Figure 9. Effect of lapatinib on ERK1/2 signaling pathways.

A. Melanoma lines expressing mutant ERBB4 exhibit increased lapatinib sensitivity with respect to ERK1 and ERK2 phosphorylation. Cells were treated for 72 hrs with 5 μ M lapatinib or vehicle as control. The activity of ERK1 and ERK2 was determined by immunoblotting with phospho-specific antibodies (α -P-ERK1/2 - recognizes phosphorylation of T202 and Y204 on ERK1 and T185 and Y187 on ERK2). Total ERK protein was also determined by immunoblotting. Shown are representative blots. B. Quantitative assessment of data from one melanoma cell line harboring WT ERBB4 and three melanoma cell lines harboring mutant ERBB4. The ratio of band intensities of P-ERK1/ERK1 or P-ERK2/ERK2 was analyzed for each melanoma cell line.

Table S1. Tyrosine Kinase genes analyzed

CCDS accession and amplifier number	Ref Seq accession and amplifier number	Gene Name	Gene Description
CCDS35165.1	NM_007313.2	ABL1/ABL	v-abl Abelson murine leukemia viral oncogene homolog 1
CCDS30947.1	NM_007314.2	ABL2/ARG	v-abl Abelson murine leukemia viral oncogene homolog 2 (arg, Abelson-related gene)
CCDS33928.1	NM_005781.4	ACK1/TNK2	tyrosine kinase, non-receptor, 2
CCDS33172.1	NM_004304.3	ALK	anaplastic lymphoma kinase (Ki-1)
CCDS12575.1	NM_021913.3	AXL	AXL receptor tyrosine kinase
CCDS5982.1	NM_001715.2	BLK	B lymphoid tyrosine kinase
CCDS14168.1	NM_203281.2	BMX	BMX non-receptor tyrosine kinase
CCDS13524.1	NM_005975.2	BRK/PTK6	PTK6 protein tyrosine kinase 6
CCDS14482.1	NM_000611.1	BTK	Bruton agammaglobulinemia tyrosine kinase
CCDS4302.1	NM_005211.2	CSF1R	colony stimulating factor 1 receptor, formerly McDonough feline sarcoma viral (v-fms) oncogene homolog
CCDS10269.1	NM_004383.1	CSK	c-src tyrosine kinase
CCDS4690.1	NM_001954.4	DDR1	discoidin domain receptor family, member 1
CCDS1241.1	NM_006182.2	DDR2	discoidin domain receptor family, member 2
CCDS5514.1	NM_005228.3	EGFR	epidermal growth factor receptor (erythroblastic leukemia viral (v-erb-b) oncogene homolog, avian)
CCDS5884.1	NM_005232.3	EPHA1	ephrin receptor EphA1
CCDS169.1	NM_004431.2	EPHA2	ephrin receptor EphA2
CCDS2922.1	NM_005233.5	EPHA3	ephrin receptor EphA3 isoform a precursor
CCDS2447.1	NM_004438.3	EPHA4	ephrin receptor EphA4
CCDS3514.1	NM_182472.1	EPHA5	ephrin receptor EphA5 isoform b
N/A	NM_001080448.2	EPHA6	EPH receptor A6 isoform a
CCDS5031.1	NM_004440.2	EPHA7	ephrin receptor EphA7
CCDS30626.1	NM_00106943.1	EPHA8	EPH receptor A8 isoform 2 precursor
CCDS425.1	NM_173641.2	EPHA10	EPH receptor A10 isoform 2
N/A	NM_004441.3	EPHB1	ephrin receptor EphB1 precursor
CCDS230.1	NM_004442.6	EPHB2	ephrin receptor EphB2 isoform 2 precursor
CCDS3268.1	NM_004443.3	EPHB3	ephrin receptor EphB3 precursor
CCDS5706.1	NM_004444.4	EPHB4	ephrin receptor EphB4 precursor
CCDS5873.1	NM_004445.2	EPHB6	ephrin receptor EphB6 precursor
CCDS32642.1	NM_004448.2	ERBB2	v-erb-b2 erythroblastic leukemia viral oncogene homolog 2, neuro/glioblastoma derived oncogene homolog (avian)
CCDS31833.1	NM_001982.2	ERBB3	v-erb-b2 erythroblastic leukemia viral oncogene homolog 3 (avian)
CCDS2394.1	NM_005235.2	ERBB4	v-erb-a erythroblastic leukemia viral oncogene homolog 4 (avian)
CCDS6381.1	NM_153831.2	FAK/PTK2	PTK2 protein tyrosine kinase 2
CCDS4098.1	NM_005246.2	FER	fer (fps/fes related) tyrosine kinase (phosphoprotein NCP94)
CCDS10365.1	NM_002005.2	FES	V-FES feline sarcoma viral/V-FPS fujinami avian
CCDS6107.1	NM_023110.2	FGFR1	fibroblast growth factor receptor 1 (fms-related tyrosine kinase 2, Pfeiffer syndrome)
CCDS31298.1	NM_000141.3	FGFR2	fibroblast growth factor receptor 2 (bacteria-expressed kinase, keratinocyte growth factor receptor)
CCDS3353.1	NM_000142.2	FGFR3	fibroblast growth factor receptor 3 (achondroplasia, thanatophoric dwarfism)
CCDS4410.1	NM_002011.3	FGFR4	fibroblast growth factor receptor 4 isoform 1
CCDS305.1	NM_005248.2	FGR	Gardher-Rasheed feline sarcoma viral (v-fgr) oncogene homolog
CCDS9330.1	NM_002019.3	FLT1/VEGFR1	fms-related tyrosine kinase 1 (vascular endothelial growth factor/vascular permeability factor receptor)
CCDS31953.1	NM_004119.2	FLT3	fms-related tyrosine kinase 3
CCDS4457.1	NM_182925.3	FLT4/VEGFR3	fms-related tyrosine kinase 4
CCDS5103.1	NM_002031.2	FRK	fyn-related kinase
CCDS5094.1	NM_002037.3	FYN	FYN oncogene related to SRC, FGR, YES
CCDS33460.1	NM_002110.2	HCK	hemopoietic cell kinase
CCDS10378.1	NM_000875.3	IGF1R	insulin-like growth factor 1 receptor
CCDS12176.1	NM_000208.2	INSR	insulin receptor
CCDS1160.1	NM_014215.1	INSR	insulin receptor-related receptor
CCDS4336.1	NM_005546.3	ITK	IL2-inducible T-cell kinase
N/A	NM_002227.2	JAK1	Janus kinase 1
CCDS6457.1	NM_004972.2	JAK2	Janus kinase 2
CCDS12366.1	NM_000215.2	JAK3	Janus kinase 3
CCDS3497.1	NM_002253.1	KDR/VEGFR2	kinase insert domain receptor (a type III receptor tyrosine kinase)
CCDS3496.1	NM_000222.2	KIT	v-kit Hardy-Zuckerman 4 feline sarcoma viral oncogene homolog
CCDS359.1	NM_005356.3	LCK	lymphocyte-specific protein tyrosine kinase
CCDS10078.1	NM_206961.1	LTK	leukocyte tyrosine kinase
CCDS6162.1	NM_002350.2	LYN	v-yes-1 Yamaguchi sarcoma viral related oncogene homolog
CCDS12113.1	NM_002378.3	MATK	megakaryocyte-associated tyrosine kinase
CCDS2094.1	NM_006343.2	MERTK/MER	c-mer proto-oncogene tyrosine kinase
N/A	NM_000245.2	MET	met proto-oncogene (hepatocyte growth factor receptor)
CCDS2807.1	NM_002447.2	MST1R/RON	macrophage stimulating 1 receptor (c-met-related tyrosine kinase)
N/A	NM_005592.1	MUSK	muscle, skeletal, receptor tyrosine kinase
CCDS1161.1	NM_002529.3	NTRK1	neurotrophic tyrosine kinase, receptor, type 1
CCDS35053.1	NM_001007097.1	NTRK2	neurotrophic tyrosine kinase receptor type 2
CCDS32322.1	NM_001012338.1	NTRK3	neurotrophic tyrosine kinase receptor type 3
CCDS3495.1	NM_006206.3	PDGFRα	platelet-derived growth factor receptor alpha
CCDS4303.1	NM_002609.3	PDGFRβ	platelet-derived growth factor receptor beta
CCDS4884.1	NM_002821.3	PTK7	PTK7 protein tyrosine kinase 7
CCDS6057.1	NM_004103.3	PYK2/PTK2B	PTK2B protein tyrosine kinase 2 beta
CCDS7200.1	NM_020975.4	RET	ret proto-oncogene
CCDS626.1	NM_005012.2	ROR1	receptor tyrosine kinase-like orphan receptor 1
CCDS6691.1	NM_004560.2	ROR2	receptor tyrosine kinase-like orphan receptor 2
CCDS5116.1	NM_002944.2	ROS1	v-ros UR2 sarcoma virus oncogene homolog 1 (avian)
N/A	NM_001005861.2	RYK	RYK receptor-like tyrosine kinase
CCDS13294.1	NM_005417.3	SRC	v-src sarcoma (Schmidt-Ruppin A-2) viral oncogene homolog (avian)
CCDS13525.1	NM_080823.2	SRMS	src-related kinase lacking C-terminal regulatory tyrosine and N-terminal myristylation sites
CCDS6688.1	NM_003177.3	SYK	spleen tyrosine kinase
CCDS3481.1	NM_003215.2	TEC	tec protein tyrosine kinase
CCDS6519.1	NM_000459.2	TEK	TEK tyrosine kinase, endothelial (venous malformations, multiple cutaneous and mucosal)
CCDS482.1	NM_005424.2	TIE	tyrosine kinase with immunoglobulin-like and EGF-like domains 1
N/A	NM_003985.3	TNK1	tyrosine kinase, non-receptor, 1
CCDS3480.1	NM_003328.2	TXK	TXK tyrosine kinase
CCDS12236.1	NM_003331.3	TYK2	tyrosine kinase 2
CCDS10080.1	NM_006293.2	TYRO3	TYRO3 protein tyrosine kinase
CCDS11824.1	NM_005433.3	YES1	v-yes-1 Yamaguchi sarcoma viral oncogene homolog 1
CCDS33254.1	NM_001079.3	ZAP70	zeta-chain (TCR) associated protein kinase 70kDa

Table S2. Primers used for PCR amplification and sequencing of kinase domains

EPHA8_Exon-10:EPHA8_Exon-11	CCDS30626_1	NM_00106943_1	AGACTCCACAGCCCCATAGA	CAGCCCTCCCGTAGGAGACATT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
EPHA8_Exon-11:EPHA8_Exon-12	CCDS30626_1	NM_00106943_1	CAGGGAGCAGGGGTGAGCC	CAGGAAGGTGTCAGAGGC	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
EPHA8_Exon-12:EPHA8_Exon-13	CCDS30626_1	NM_00106943_1	CATCGCTCTATCCATCTCG	CAATGACTGCTTACCCCATTTG	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
EPHA8_Exon-13:EPHA8_Exon-14	CCDS30626_1	NM_00106943_1	CTCGCTCTATCCATCTCG	CTCGCTCTATCCATCTCG	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
EPHA8_Exon-14:EPHA8_Exon-15	CCDS30626_1	NM_00106943_1	GTCCCTCTGACTCTGAAACA	TAGTGTCAGGCCAACAGTC	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
EPHA10_Exon10	CCDS31305_1	NM_00109943_9	CTTCCAGACAGCACAGAGAC	TGGTCTAGGGCTACCGGTGTTAT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
EPHA10_Exon13	CCDS31305_1	NM_00109943_9	GTGAGGGAGGAAGACTGTGAG	GCCCAGGGATGCCCTTCTGA	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
EPHA10_Exon13:EPHA10_Exon14	CCDS31305_1	NM_00109943_9	CCCAAATCTCAGCTTACATCT	GATGATGTCAGGAGTACATGA	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
EPHA10_Exon15	CCDS31305_1	NM_00109943_9	CAGAACAGAACAGAACAGAG	CTGGCTGAGGGCTTAACT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
EPHA11_Exon10	CCDS31305_1	NM_00109943_9	AAGCTTCTTCTTCTTCTTCT	DATACTTCTTCTTCTTCTTCT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
EPHB1_Exon-11	N/A	N/A	GGCAACTTAATACAAATTCAC	GCACAAATCTCTGAACTGTC	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
EPHB1_Exon-12	N/A	N/A	AGTGGGAAGAGGAGGAGAC	CTCACAGTCAAAGGCAAAT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
EPHB1_Exon-13	N/A	N/A	NTCACTCTGAGCTCTCTAC	TACTGCTTAAATTGGGGCTGT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
EPHB1_Exon-14	N/A	N/A	TCGACAGAAATTCATTAACAGC	CCAGGTATCAAAAGGAAATC	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
EPHB1_Exon-15	CCDS32301_1	NM_0044413	CAGGGAGATGGCCAGGTCTTAT	GTGCAAGATGGCCAGGTCTTAT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
EPHB1_Exon-16	CCDS32301_1	NM_0044413	QAATGATGGCTTACCTTCTC	AGATGATGGCTTACCTTCTC	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
EPHB2_Exon-12	CCDS32301_1	NM_0044426	GTCCAAACATCTGAGCTCT	GTCAAGGAGCTTCTGCACTAG	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
EPHB2_Exon-13	CCDS32301_1	NM_0044426	TTGGAGATGGAAAGGAGGTTTC	AATAATGCTGTAAGGAGGTTTC	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
EPHB2_Exon-14	CCDS32301_1	NM_0044426	CTAGATGTTGDTGAAACCGAGTC	TTAAAAGATGCTCTCTGAG	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
EPHB3_Exon-10:EPHB3_Exon-11	CCDS32634_1	NM_004443_3	GATTAGGGCAGGACAAAGAG	TATGGACTCTGAGTGGGCGGTG	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
EPHB3_Exon-12:EPHB3_Exon-13	CCDS32634_1	NM_004443_3	TTGGCTGAGCTTCTGAGAAGC	TGTCAGACATCTGAGTATG	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
EPHB3_Exon-13:EPHB3_Exon-14	CCDS32634_1	NM_004443_3	CTGCTGCTTACAGGAGGAAAG	TTGCTGCTTACAGGAGGAAAG	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
EPHB4_Exon-14	CCDS57061_1	NM_004444_4	CAAGAACCCAGCAGTGTGAG	AACCAAGGATGGACAGGGGT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
EPHB4_Exon-15	CCDS57061_1	NM_004444_4	ACACCAAACTGACAGCTCTT	CTATGAGTGGCCACTGTACTGC	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
EPHB4_Exon-15:EPHB4_Exon-16	CCDS57061_1	NM_004444_4	AGTGCAGAAAGAGCACAGCTTG	ATATACTCTGCTGAAAGAC	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
EPHB4_Exon-16:EPHB4_Exon-17	CCDS58731_1	NM_004444_4	TTGGGGATGGAAAGGAGGTTTC	CTTCGGCTTCTTCCAGGTAT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
EPHB4_Exon-17	CCDS58731_1	NM_004444_4	GGATGTTGAGTGGAAAGGTTTC	GATGATGGGGCTGGCTGGAGA	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
EPHB4_Exon-18:EPHB4_Exon-19	CCDS58731_1	NM_004445_2	GTGCTGCTTACAGGAGGAAAG	CTTCTGCTGCTGCTGAAAT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
EPHB4_Exon-19	CCDS58731_1	NM_004445_2	GTGCTGCTTACAGGAGGAAAG	TGTTGCTTACAGGAGGAAAG	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
ERBB2_Exon-21:ERBB2_Exon-22:ERBB2_Exon-23	CCDS2642_1	NM_004448_2	GGGGCAAGTGGATGAGGAGAC	ATACATGCTGGCTCCCTTCT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
ERBB2_Exon-22	CCDS2642_1	NM_004448_2	GGGGCAAGTGGATGAGGAGAC	GATGATGGGGCTGGCTGGAG	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
ERBB2_Exon-23	CCDS2642_1	NM_004448_2	CTGGCTGAGAACGGGGAGAT	ACAGAGTGGCTTCTCAGAGT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
ERBB2_Exon-23:ERBB2_Exon-24	CCDS2642_1	NM_004448_2	CTGGCTGAGAACGGGGAGAT	TGTTGCTTACAGGAGGAAAT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
ERBB2_Exon-24	CCDS2642_1	NM_004448_2	CTGGCTGAGAACGGGGAGAT	CTGGCTGAGAACGGGGAGAT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
ERBB2_Exon-19:ERBB2_Exon-20	CCDS1833_1	NM_001982_2	TAAGACCAAACTCCCTTGTGAT	TTTCTGCTGCTGCTGCTGAA	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
ERBB2_Exon-20:ERBB2_Exon-21	CCDS1833_1	NM_001982_2	TAAGACCAAACTCCCTTGTGAT	AAAGAGAAAAATTGTGGGCTCAG	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
ERBB2_Exon-21:ERBB2_Exon-22	CCDS1833_1	NM_001982_2	TAAGACCAAACTCCCTTGTGAT	AAAGAGAAAAATTGTGGGCTCAG	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
ERBB2_Exon-22:ERBB2_Exon-23	CCDS1833_1	NM_001982_2	TAAGACCAAACTCCCTTGTGAT	AAAGAGAAAAATTGTGGGCTCAG	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
ERBB3_Exon-24	CCDS1833_1	NM_001982_2	TAAGACCAAACTCCCTTGTGAT	AAAGAGAAAAATTGTGGGCTCAG	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
ERBB4_Exon-18	CCDS2394_1	NM_005238_2	TCATTGTTGCTGAGCACTCTG	CTGCTGCTTACCTCATCTTCT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
ERBB4_Exon-19	CCDS2394_1	NM_005238_2	TCATTGTTGCTGAGCACTCTG	CTGCTGCTTACCTCATCTTCT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
ERBB4_Exon-20	CCDS2394_1	NM_005238_2	TCATTGTTGCTGAGCACTCTG	CTGCTGCTTACCTCATCTTCT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
ERBB4_Exon-21	CCDS2394_1	NM_005238_2	TCATTGTTGCTGAGCACTCTG	CTGCTGCTTACCTCATCTTCT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
ERBB4_Exon-22	CCDS2394_1	NM_005238_2	TCATTGTTGCTGAGCACTCTG	CTGCTGCTTACCTCATCTTCT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
ERBB4_Exon-23	CCDS2394_1	NM_005238_2	TCATTGTTGCTGAGCACTCTG	CTGCTGCTTACCTCATCTTCT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
ERBB4_Exon-24	CCDS2394_1	NM_005238_2	TCATTGTTGCTGAGCACTCTG	CTGCTGCTTACCTCATCTTCT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
ERBB4_Exon-25	CCDS2394_1	NM_005238_2	TCATTGTTGCTGAGCACTCTG	CTGCTGCTTACCTCATCTTCT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FER_Exon-13	CCDS1833_1	NM_001982_2	ACCTCTTCTCTGCTTACATTC	TGCTGCTTACCTCATCTTCT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FER_Exon-14	CCDS1833_1	NM_001982_2	ACCTCTTCTCTGCTTACATTC	AACTCTTCTCTGCTTACATTC	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FER_Exon-15	CCDS1833_1	NM_001982_2	ACCTCTTCTCTGCTTACATTC	AACTCTTCTCTGCTTACATTC	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FER_Exon-16	CCDS1833_1	NM_001982_2	ACCTCTTCTCTGCTTACATTC	AACTCTTCTCTGCTTACATTC	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FES_Exon-13:FES_Exon-14:FES_Exon-15	CCDS10365_1	NM_002008_2	ACCGAGGCTGGCTGGAAATAC	TGCTGCTTACCTCATCTTCT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FES_Exon-15:FES_Exon-16:FES_Exon-17	CCDS10365_1	NM_002008_2	ACCGAGGCTGGCTGGAAATAC	TGCTGCTTACCTCATCTTCT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FES_Exon-16	CCDS10365_1	NM_002008_2	ACCGAGGCTGGCTGGAAATAC	TGCTGCTTACCTCATCTTCT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FES_Exon-17	CCDS10365_1	NM_002008_2	ACCGAGGCTGGCTGGAAATAC	TGCTGCTTACCTCATCTTCT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FES_Exon-18	CCDS10365_1	NM_002008_2	ACCGAGGCTGGCTGGAAATAC	TGCTGCTTACCTCATCTTCT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGFR1_Exon-11:FGFR1_Exon-12	CCDS1071_1	NM_023110_2	AAACAGTACAGCAGGCTAC	GTACATGCTTACCTGAGCTGT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGFR1_Exon-12:FGFR1_Exon-13	CCDS1071_1	NM_023110_2	AAACAGTACAGCAGGCTAC	GTACATGCTTACCTGAGCTGT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGFR1_Exon-13:FGFR1_Exon-14:FGFR1_Exon-15	CCDS1071_1	NM_023110_2	AAACAGTACAGCAGGCTAC	GTACATGCTTACCTGAGCTGT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGFR1_Exon-14:FGFR1_Exon-16	CCDS1071_1	NM_023110_2	AAACAGTACAGCAGGCTAC	GTACATGCTTACCTGAGCTGT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGFR2_Exon-10	CCDS3298_1	NM_0010941_3	TCATTGTTGCTGGCTGGCTAT	TGCTGCTTACCTCATCTTCT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGFR2_Exon-11	CCDS3298_1	NM_0010941_3	TCATTGTTGCTGGCTGGCTAT	AAACAGCTTACATCTTCT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGFR2_Exon-12	CCDS3298_1	NM_0010941_3	TCATTGTTGCTGGCTGGCTAT	AAACAGCTTACATCTTCT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGFR2_Exon-13	CCDS3298_1	NM_0010941_3	TCATTGTTGCTGGCTGGCTAT	AAACAGCTTACATCTTCT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGFR2_Exon-14	CCDS3298_1	NM_0010941_3	TCATTGTTGCTGGCTGGCTAT	AAACAGCTTACATCTTCT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGFR2_Exon-15	CCDS3298_1	NM_0010941_3	TCATTGTTGCTGGCTGGCTAT	AAACAGCTTACATCTTCT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGFR2_Exon-16	CCDS3298_1	NM_0010941_3	TCATTGTTGCTGGCTGGCTAT	AAACAGCTTACATCTTCT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGFR2_Exon-17	CCDS3298_1	NM_0010941_3	TCATTGTTGCTGGCTGGCTAT	AAACAGCTTACATCTTCT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGFR2_Exon-18	CCDS3298_1	NM_0010941_3	TCATTGTTGCTGGCTGGCTAT	AAACAGCTTACATCTTCT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGFR3_Exon-10:FGFR3_Exon-11:FGFR3_Exon-12	CCDS3335_1	NM_0010414_3	TAATGGTTTCTGGCTGGCTAT	TGCTGCTTACCTCATCTTCT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGFR3_Exon-11:FGFR3_Exon-12:FGFR3_Exon-13	CCDS3335_1	NM_0010414_3	TAATGGTTTCTGGCTGGCTAT	AAACACAGTACAGCTTAC	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGFR3_Exon-12:FGFR3_Exon-13:FGFR3_Exon-14	CCDS3335_1	NM_0010414_3	TAATGGTTTCTGGCTGGCTAT	AAACACAGTACAGCTTAC	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGFR3_Exon-13:FGFR3_Exon-14:FGFR3_Exon-15	CCDS3335_1	NM_0010414_3	TAATGGTTTCTGGCTGGCTAT	AAACACAGTACAGCTTAC	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGFR3_Exon-14:FGFR3_Exon-16	CCDS3335_1	NM_0010414_3	TAATGGTTTCTGGCTGGCTAT	AAACACAGTACAGCTTAC	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGFR3_Exon-15:FGFR3_Exon-16	CCDS3335_1	NM_0010414_3	TAATGGTTTCTGGCTGGCTAT	AAACACAGTACAGCTTAC	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGFR3_Exon-16:FGFR3_Exon-17	CCDS3335_1	NM_0010414_3	TAATGGTTTCTGGCTGGCTAT	AAACACAGTACAGCTTAC	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGFR3_Exon-17:FGFR3_Exon-18	CCDS3335_1	NM_0010414_3	TAATGGTTTCTGGCTGGCTAT	AAACACAGTACAGCTTAC	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGFR4_Exon-10	CCDS4410_1	NM_002011_3	CACTGAGTCTCCGACCTCTAC	TGCTGCTTACCTCATCTTCT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGFR4_Exon-11	CCDS4410_1	NM_002011_3	CACTGAGTCTCCGACCTCTAC	AAAGAGCTTACATCTTCT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGFR4_Exon-12	CCDS4410_1	NM_002011_3	CACTGAGTCTCCGACCTCTAC	AAAGAGCTTACATCTTCT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGFR4_Exon-13	CCDS4410_1	NM_002011_3	CACTGAGTCTCCGACCTCTAC	AAAGAGCTTACATCTTCT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGFR4_Exon-14	CCDS4410_1	NM_002011_3	CACTGAGTCTCCGACCTCTAC	AAAGAGCTTACATCTTCT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGFR4_Exon-15	CCDS4410_1	NM_002011_3	CACTGAGTCTCCGACCTCTAC	AAAGAGCTTACATCTTCT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGFR4_Exon-16	CCDS4410_1	NM_002011_3	CACTGAGTCTCCGACCTCTAC	AAAGAGCTTACATCTTCT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGFR4_Exon-17	CCDS4410_1	NM_002011_3	CACTGAGTCTCCGACCTCTAC	AAAGAGCTTACATCTTCT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGFR4_Exon-18	CCDS4410_1	NM_002011_3	CACTGAGTCTCCGACCTCTAC	AAAGAGCTTACATCTTCT	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGR_Exon-10	CCDS3930_1	NM_002019_3	TTGGCTGAGGAGGAGGAGGAG	TGAGGAGGAGGAGGAGGAG	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGR_Exon-11	CCDS3930_1	NM_002019_3	TTGGCTGAGGAGGAGGAGGAG	TGAGGAGGAGGAGGAGGAG	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGR_Exon-12	CCDS3930_1	NM_002019_3	TTGGCTGAGGAGGAGGAGGAG	TGAGGAGGAGGAGGAGGAG	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGR_Exon-13	CCDS3930_1	NM_002019_3	TTGGCTGAGGAGGAGGAGGAG	TGAGGAGGAGGAGGAGGAG	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGR_Exon-14	CCDS3930_1	NM_002019_3	TTGGCTGAGGAGGAGGAGGAG	TGAGGAGGAGGAGGAGGAG	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGR_Exon-15	CCDS3930_1	NM_002019_3	TTGGCTGAGGAGGAGGAGGAG	TGAGGAGGAGGAGGAGGAG	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGR_Exon-16	CCDS3930_1	NM_002019_3	TTGGCTGAGGAGGAGGAGGAG	TGAGGAGGAGGAGGAGGAG	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGR_Exon-17	CCDS3930_1	NM_002019_3	TTGGCTGAGGAGGAGGAGGAG	TGAGGAGGAGGAGGAGGAG	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGR_Exon-18	CCDS3930_1	NM_002019_3	TTGGCTGAGGAGGAGGAGGAG	TGAGGAGGAGGAGGAGGAG	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGR_Exon-19	CCDS3930_1	NM_002019_3	TTGGCTGAGGAGGAGGAGGAG	TGAGGAGGAGGAGGAGGAG	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGR_Exon-20	CCDS3930_1	NM_002019_3	TTGGCTGAGGAGGAGGAGGAG	TGAGGAGGAGGAGGAGGAG	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGR_Exon-21	CCDS3930_1	NM_002019_3	TTGGCTGAGGAGGAGGAGGAG	TGAGGAGGAGGAGGAGGAG	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGR_Exon-22	CCDS3930_1	NM_002019_3	TTGGCTGAGGAGGAGGAGGAG	TGAGGAGGAGGAGGAGGAG	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGR_Exon-23	CCDS3930_1	NM_002019_3	TTGGCTGAGGAGGAGGAGGAG	TGAGGAGGAGGAGGAGGAG	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGR_Exon-24	CCDS3930_1	NM_002019_3	TTGGCTGAGGAGGAGGAGGAG	TGAGGAGGAGGAGGAGGAG	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGR_Exon-25	CCDS3930_1	NM_002019_3	TTGGCTGAGGAGGAGGAGGAG	TGAGGAGGAGGAGGAGGAG	TGTAAAACCGACGCCAGT	CAGGAACACGCTATGACC
FGR_Exon-26	CCDS3930_1	NM_002019_3	TTGGCTGAGGAGGAGGAGGAG	TGAGGAGGAGGAGGAGGAG	TGTAAAACCGACGCCAGT	CAGGA

TNK1_Exon-6	N/A	NM_003985.3	AAAGAGGACCTTTGTGAAAGATGG	TGTCAACACGACTGCAACTG	TGTAAAACCGACGGCCAGT	CAGGAACACGCTATGACC
TXK_Exon-10	CCDS3480.1	NM_003328.2	GACATTGAGCTTGAGAAGGT	TCCCAGCTAGTTGGTAGG	TGTAAAACCGACGGCCAGT	CAGGAACACGCTATGACC
TXK_Exon-11	CCDS3480.1	NM_003328.2	TAATTTCGACCATACTACCAA	AATCACTCTTCCCTCCCTTG	TGTAAAACCGACGGCCAGT	CAGGAACACGCTATGACC
TXK_Exon-13	CCDS3480.1	NM_003328.2	ADAACTGCTTCTTCTTAA	GGAACTTCTTCTTCTTCTTG	TGTAAAACCGACGGCCAGT	CAGGAACACGCTATGACC
TXK_Exon-15	CCDS3480.1	NM_003328.2	TCCAAAGAACATCTCAAACA	AAAAAGAAAAGAAATCTACTGG	TGTAAAACCGACGGCCAGT	CAGGAACACGCTATGACC
TYK2_Exon-10;TYK2_Exon-11	CCDS12236.1	NM_003313.3	GTGCAAATCCCTCACACATA	TCACGCCCTACTCAACTG	TGTAAAACCGACGGCCAGT	CAGGAACACGCTATGACC
TYK2_Exon-10;TYK2_Exon-11;TYK2_Exon-12	CCDS12236.1	NM_003313.3	GCTCTCTGCTCATCATCTTG	CTACGCCCTGATCTCACAG	TGTAAAACCGACGGCCAGT	CAGGAACACGCTATGACC
TYK2_Exon-10;TYK2_Exon-11;TYK2_Exon-12	CCDS12236.1	NM_003313.3	AGGCTGGCTGCTCTCGTAGAA	GGGTGACCAAGAGGAGATCA	TGTAAAACCGACGGCCAGT	CAGGAACACGCTATGACC
TYK2_Exon-14;TYK2_Exon-15	CCDS12236.1	NM_003313.3	TCAGCTGATACCCAGAGAGA	AATGTCACAGTTGCTTTTCA	TGTAAAACCGACGGCCAGT	CAGGAACACGCTATGACC
TYK2_Exon-14;TYK2_Exon-15	CCDS12236.1	NM_003313.3	CTGCTGATACCCAGAGAGA	ATAGTCACAGTTGCTTTTCA	TGTAAAACCGACGGCCAGT	CAGGAACACGCTATGACC
TYK2_Exon-16	CCDS12236.1	NM_003313.3	CTTATGAACTCCACTGCAAGAA	GCCCCAGAAGAGGTTTATCA	TGTAAAACCGACGGCCAGT	CAGGAACACGCTATGACC
TYK2_Exon-17;TYK2_Exon-18	CCDS12236.1	NM_003313.3	GTACTGCAGCTGCTGGGTT	CTGCTGGGAAAGGATCG	TGTAAAACCGACGGCCAGT	CAGGAACACGCTATGACC
TYK2_Exon-17;TYK2_Exon-18	CCDS12236.1	NM_003313.3	CTGCTCTGAGGGCTTTCAC	CCTCCAAAACAGGCTGAGT	TGTAAAACCGACGGCCAGT	CAGGAACACGCTATGACC
TYK2_Exon-18;TYK2_Exon-19	CCDS12236.1	NM_003313.3	GTTCCGAGAGTCACTGCAAG	CACCGACACATCATCAAGTACA	TGTAAAACCGACGGCCAGT	CAGGAACACGCTATGACC
TYK2_Exon-19;TYK2_Exon-20	CCDS12236.1	NM_003313.3	CTTGTGCTGATACCCAGAT	GGCTTCTTCTTCTTCTTCT	TGTAAAACCGACGGCCAGT	CAGGAACACGCTATGACC
TYK2_Exon-20;TYK2_Exon-21	CCDS12236.1	NM_003313.3	ACCCGAAAGGACAGACAT	CCCCCCACTGAACTCTAC	TGTAAAACCGACGGCCAGT	CAGGAACACGCTATGACC
TYK2_Exon-22	CCDS12236.1	NM_003313.3	CCTCTCCACAGCAGGATGT	CCAAACACTGACTCACGCTTA	TGTAAAACCGACGGCCAGT	CAGGAACACGCTATGACC
TYK2_Exon-22;TYK2_Exon-23	CCDS12236.1	NM_003313.3	AGACAGGAGTAAGGCAACAC	CTGCCCTTCTTGCCTCTTG	TGTAAAACCGACGGCCAGT	CAGGAACACGCTATGACC
TYR03_Exon-12;TYR03_Exon-13	CCDS10080.1	NM_006293.2	ACAAATCTGCCCTGTGGAGGT	AGGCTACAGCTATCCCCACT	TGTAAAACCGACGGCCAGT	CAGGAACACGCTATGACC
TYR03_Exon-13	CCDS10080.1	NM_006293.2	GGAACTTCTGCTGATAGGT	AGCTTACGCTTACGATGTC	TGTAAAACCGACGGCCAGT	CAGGAACACGCTATGACC
TYR03_Exon-14	CCDS10080.1	NM_006293.2	TTACAGCTGATACACAGCA	TCTTCATGCTACAGTC	TGTAAAACCGACGGCCAGT	CAGGAACACGCTATGACC
TYR03_Exon-15;TYR03_Exon-16	CCDS10080.1	NM_006293.2	GCTACTCCAGACGACAGTGT	CTGAGCGGCTGAGTTCTACTG	TGTAAAACCGACGGCCAGT	CAGGAACACGCTATGACC
TYR03_Exon-16;TYR03_Exon-17	CCDS10080.1	NM_006293.2	GTCCTCCAGGCTCCCTTCA	CTGTCACACACTCTACTG	TGTAAAACCGACGGCCAGT	CAGGAACACGCTATGACC
TYR03_Exon-17;TYR03_Exon-18	CCDS10080.1	NM_006293.2	GGTACTCTGGAGGAAAGATA	CCATCAGCCAAACTCCAGAT	TGTAAAACCGACGGCCAGT	CAGGAACACGCTATGACC
TYR03_Exon-18	CCDS10080.1	NM_006293.2	CAGGACACTCTAGTCACTCT	CCATCAGCTTCTCCAGAGT	TGTAAAACCGACGGCCAGT	CAGGAACACGCTATGACC
TYR03_Exon-19	CCDS10080.1	NM_006293.2	AACCTCTGGCTCAAATAATACC	CAGITCCCTTCTCCAGAGT	TGTAAAACCGACGGCCAGT	CAGGAACACGCTATGACC
YES1_Exon-10	CCDS11824.1	NM_006433.3	TATGCTGCTGATACCTCTT	GCACAGCTGGAGAGCTAC	TGTAAAACCGACGGCCAGT	CAGGAACACGCTATGACC
YES1_Exon-11	CCDS11824.1	NM_006433.3	AGGCTGCTGATACCTCTT	AGGCTGCTGATACCTCTT	TGTAAAACCGACGGCCAGT	CAGGAACACGCTATGACC
YES1_Exon-6	CCDS11824.1	NM_005433.3	TTTTAAAGATTGAGACCAAACCTCA	TAATCAATGGGGTGGCTT	TGTAAAACCGACGGCCAGT	CAGGAACACGCTATGACC
YES1_Exon-6;YES1_Exon-7	CCDS11824.1	NM_005433.3	ACAAAGTGTAGTTCTCTATCC	CCATCTGCCCTACGTTCTTA	TGTAAAACCGACGGCCAGT	CAGGAACACGCTATGACC
YES1_Exon-8	CCDS11824.1	NM_005433.3	TCCCCACATTCTACACTACATC	AGAACATGGCTGATGGTTATGG	TGTAAAACCGACGGCCAGT	CAGGAACACGCTATGACC
YES1_Exon-9	CCDS11824.1	NM_005433.3	CGCTTAATCAGCTTCTACATAC	CCGTAATCAGACTCTAGCCCTT	TGTAAAACCGACGGCCAGT	CAGGAACACGCTATGACC
ZAP70_Exon-12	CCDS3254.1	NM_001079.3	AGCTTACGCTGATACGTTCT	AAATTACGCTGATAGGTGG	TGTAAAACCGACGGCCAGT	CAGGAACACGCTATGACC
ZAP70_Exon-7	CCDS3254.1	NM_001079.3	CTGTCAGGAAACGGCATGG	CTGTCAGGAAACGGCATGG	TGTAAAACCGACGGCCAGT	CAGGAACACGCTATGACC
ZAP70_Exon-8	CCDS3254.1	NM_001079.3	CCTGGGTGCGACTACAGTT	ATCCCGAAAAGATCCAAAAGT	TGTAAAACCGACGGCCAGT	CAGGAACACGCTATGACC
ZAP70_Exon-9;ZAP70_Exon-10	CCDS3254.1	NM_001079.3	GTGTTGGTGTGCTCTAAAGC	GTGACCCCATGCTCCAGAC	TGTAAAACCGACGGCCAGT	CAGGAACACGCTATGACC
ZAP70_Exon-9;ZAP70_Exon-10;ZAP70_Exon-11	CCDS3254.1	NM_001079.3	CTACGGCCAGATCAGGACCTT	ACTGGAGCAGACAGCAAC	TGTAAAACCGACGGCCAGT	CAGGAACACGCTATGACC

Table S3. Primers used for PCR amplification and sequencing of the whole coding region

FER-1	CCDS4098.1	NM_005246.1	GTAaaaACGCGGCCAGTGTCTGACTTAAGATGCTCATTAATAGTT	TTCATCTGTTCTGATCTTGG	GTAaaaACGCGGCCAGT
FER-2	CCDS4098.1	NM_005246.1	GTAaaaACGCGGCCAGTTCTGTGTTGGAAATGATGAA	TTGAAACAGTTCGCCAGAACCT	GTAaaaACGCGGCCAGT
FER-3	CCDS4098.1	NM_005246.1	TGACAATTAGCCAATGTGGGT	GTAaaaACGCGGCCAGTCTGGCTGATTTCT	GTAaaaACGCGGCCAGT
FER-4	CCDS4098.1	NM_005246.1	GTAaaaACGCGGCCAGTAGAACTGATTAATGTTAGGTTAAACCA	TTTCAACTATCAAGCTTCTG	GTAaaaACGCGGCCAGT
FER-5	CCDS4098.1	NM_005246.1	GTAaaaACGCGGCCAGTAGGCTTAAATGTTAGGTTAAACCA	AGGTGTTACAGGAAAACCCA	GTAaaaACGCGGCCAGT
FER-6	CCDS4098.1	NM_005246.1	GTAaaaACGCGGCCAGTACACAGAACCTTGATATTCTGG	TTTGAACCCAGGATTTAGTTTG	GTAaaaACGCGGCCAGT
FER-7	CCDS4098.1	NM_005246.1	GTAaaaACGCGGCCAGTAGAAATGTTAGGTCAACTGCC	CCACCATCTAGTAAAGTTGCC	GTAaaaACGCGGCCAGT
FER-8	CCDS4098.1	NM_005246.1	GTAaaaACGCGGCCAGTAGGCTGAGCTGACAAAG	GTTGATTGTACTCCAAACACTTC	GTAaaaACGCGGCCAGT
FER-9	CCDS4098.1	NM_005246.1	TTCTGTCTCCATTGTCATGTAC	GTAAAACGCGGCCAGTGTGATAAAGTGTGATAAGTACCC	GTAaaaACGCGGCCAGT
FER-10	CCDS4098.1	NM_005246.1	GTAaaaACGCGGCCAGTCAGCCTATTACAAACCCATTC	GGGAGGAAACATTGTAAAGAC	GTAaaaACGCGGCCAGT
FER-11	CCDS4098.1	NM_005246.1	GTAaaaACGCGGCCAGTCAAGGAACTTGTGTTCAA	TGCAATTCTAGCTGTGTTGTT	GTAaaaACGCGGCCAGT
FER-12	CCDS4098.1	NM_005246.1	AGACTTACACATCTATTCTAGTGA	GTAaaaACGCGGCCAGTCAGCTGCTCA	GTAaaaACGCGGCCAGT
FER-13	CCDS4098.1	NM_005246.1	GTAaaaACGCGGCCAGTCTGGCTTGTGATATTCTG	TAAGCACCTCTCTTCTTGG	GTAaaaACGCGGCCAGT
FER-14	CCDS4098.1	NM_005246.1	TCTTGTGAGACTTCTGAGGAGG	GTAaaaACGCGGCCAGTCTACTGTTACTGTTG	GTAaaaACGCGGCCAGT
FER-15	CCDS4098.1	NM_005246.1	CAACTGAGAACGGACACTTACAC	GTAAAACGCGGCCAGTCAAGTAAACATGGG	GTAaaaACGCGGCCAGT
FER-16	CCDS4098.1	NM_005246.1	TTTGTGTTACCTCCAAATCATC	GTAAAACGCGGCCAGTTGAGACTTACAC	GTAaaaACGCGGCCAGT
FER-17	CCDS4098.1	NM_005246.1	CCCTCTCAGCAGCACAC	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAaaaACGCGGCCAGT
FER-18	CCDS4098.1	NM_005246.1	AAGAAAGCCCTTGTAGGACTG	GTAAAACGCGGCCAGTACTCATTAATGCCCTGTG	GTAaaaACGCGGCCAGT
FLT1-1	CCDS9330.1	NM_002019.2	GTAAAACGCGGCCAGTAAAGTGTGCTCTGGCT	ACTAGTCCAGGCTCTGCT	GTAaaaACGCGGCCAGT
FLT1-2	CCDS9330.1	NM_002019.2	TTTATGAGCAAGGAGTAGCTAGAA	GTAAAACGCGGCCAGTGTGCTTCATTCGATCAC	GTAaaaACGCGGCCAGT
FLT1-3	CCDS9330.1	NM_002019.2	GTAAAACGCGGCCAGTCAAGGTTAAATGTTGCAAC	AGGAAGTACGCGGCCAGT	GTAaaaACGCGGCCAGT
FLT1-4	CCDS9330.4	NM_002019.2	GTAAAACGCGGCCAGTCAACACTGTGTTAACAGT	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAaaaACGCGGCCAGT
FLT1-5	CCDS9330.1	NM_002019.2	GTAAAACGCGGCCAGTCAAGGAGGAACTTGTG	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAaaaACGCGGCCAGT
FLT1-6	CCDS9330.6	NM_002019.2	GCATGACATACAGCGGAAGTC	GTAAAACGCGGCCAGTACTCATTAATGCCCTGTG	GTAaaaACGCGGCCAGT
FLT1-7	CCDS9330.7	NM_002019.2	CTCTAGGTGATTGACATGATTC	GTAAAACGCGGCCAGTACCATGGCCAACTGTATTC	GTAaaaACGCGGCCAGT
FLT1-8	CCDS9330.8	NM_002019.2	GTAAAACGCGGCCAGTAACTACCCCTCCGAATG	TGCTGAGGAACCTCTGCT	GTAaaaACGCGGCCAGT
FLT1-9	CCDS9330.9	NM_002019.2	GTTAAACGCGGCCAGTAAAGGTTAAATGTTGCAAC	ATGCCACATCTGGAAATACAC	GTAaaaACGCGGCCAGT
FLT1-10	CCDS9330.10	NM_002019.2	GTAAAACGCGGCCAGTAAAGCTGAGATGTTGTC	ACATCTGGACATCTGGAAATACAC	GTAaaaACGCGGCCAGT
FLT1-11	CCDS9330.11	NM_002019.2	TTTGTATCTGAGGACTTACACAA	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAaaaACGCGGCCAGT
FLT1-12	CCDS9330.12	NM_002019.2	GTAAAACGCGGCCAGTAACTCAGAAGTGTG	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAaaaACGCGGCCAGT
FLT1-13	CCDS9330.13	NM_002019.2	GTAAAACGCGGCCAGTAACTGAGATGATGAGA	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAaaaACGCGGCCAGT
FLT1-14	CCDS9330.14	NM_002019.2	GTAAAACGCGGCCAGTCAAGGAGTAAAGTGTGAAAG	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAaaaACGCGGCCAGT
FLT1-15	CCDS9330.15	NM_002019.2	GAGCTGGATTCTCTGCT	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAaaaACGCGGCCAGT
FLT1-16	CCDS9330.16	NM_002019.2	AACATTITAGGAAACTGAGCC	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAaaaACGCGGCCAGT
FLT1-17	CCDS9330.17	NM_002019.2	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAaaaACGCGGCCAGT
FLT1-18	CCDS9330.18	NM_002019.2	ACTGGAGATCTGCTCATC	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAaaaACGCGGCCAGT
FLT1-19	CCDS9330.19	NM_002019.2	GGGAAGTGTAGTGAAGC	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAaaaACGCGGCCAGT
FLT1-20	CCDS9330.20	NM_002019.2	GTAAAACGCGGCCAGTAAAGAACCTTGGCATC	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAaaaACGCGGCCAGT
FLT1-21	CCDS9330.21	NM_002019.2	TACCTTCCAGCTTCCCAGG	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAaaaACGCGGCCAGT
FLT1-22	CCDS9330.22	NM_002019.2	ATTAGGACTCTGCTGTTG	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAaaaACGCGGCCAGT
FLT1-23	CCDS9330.23	NM_002019.2	CGCTCTGCTGAGTTGTTG	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAaaaACGCGGCCAGT
FLT1-24	CCDS9330.24	NM_002019.2	GGTGGACCCAGTACTCTG	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAaaaACGCGGCCAGT
FLT1-25	CCDS9330.25	NM_002019.2	GTAAAACGCGGCCAGTAACTGGCTTGTGACCTC	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAaaaACGCGGCCAGT
FLT1-26	CCDS9330.26	NM_002019.2	GAGCTCTGTTGACCAATT	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAaaaACGCGGCCAGT
FLT1-27	CCDS9330.1	NM_002019.2	ATGGGCTATCTGGTGTGAT	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAaaaACGCGGCCAGT
FLT1-28	CCDS9330.21	NM_002019.2	ATTTCAGGACCTCTGAACTCC	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAaaaACGCGGCCAGT
FLT1-29	CCDS9330.3	NM_002019.2	AACCTCCGGACCTACAGCTG	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAaaaACGCGGCCAGT
FLT1-30	CCDS9330.4	NM_002019.2	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAaaaACGCGGCCAGT
MATK-1	CCDS1211.3	NM_002378.2	AAACACCGCTTCTTAACTGGCC	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAaaaACGCGGCCAGT
MATK-1	CCDS1211.3	NM_139355.1	GTAAAACGCGGCCAGTGGAAATGCTACTCC	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAaaaACGCGGCCAGT
MATK-2	CCDS1211.3	NM_002378.2	GTAAAACGCGGCCAGTGGAAATGCTACTCC	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAaaaACGCGGCCAGT
MATK-3	CCDS1211.3	NM_002378.2	CATCTGCTTCTCTGACCTTC	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAaaaACGCGGCCAGT
MATK-4	CCDS1211.3	NM_002378.2	GTAAAACGCGGCCAGTAACTCAGGAAACAC	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAaaaACGCGGCCAGT
MATK-5	CCDS1211.3	NM_002378.2	CAAGCTCAGCTCTG	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAaaaACGCGGCCAGT
MATK-6	CCDS1211.3	NM_002378.2	GTAAAACGCGGCCAGTAAACCGCTTCTC	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAaaaACGCGGCCAGT
MATK-7	CCDS1211.3	NM_002378.2	TTTACACCATCTCCCTCTG	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAaaaACGCGGCCAGT
MATK-8	CCDS1211.3	NM_002378.2	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAaaaACGCGGCCAGT
MATK-9	CCDS1211.3	NM_002378.2	GTGCGCGTGTGAAATATCAAG	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAaaaACGCGGCCAGT
MATK-10	CCDS1211.3	NM_002378.2	AGGAAGTACCAACAGAAC	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAaaaACGCGGCCAGT
MATK-11	CCDS1211.3	NM_002378.2	GTAAAACGCGGCCAGTGGCTTGTGACCTTC	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAaaaACGCGGCCAGT
MATK-12	CCDS1211.3	NM_002378.2	GTAAAACGCGGCCAGTGGCTTGTGACCTTC	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAaaaACGCGGCCAGT
MATK-13	CCDS1211.3	NM_002378.2	GTAAAACGCGGCCAGTGGCTTGTGACCTTC	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAaaaACGCGGCCAGT
MET-1	N/A	NM_002245		GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGT
MET-1	N/A	NM_002245		GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGT
MET-1	N/A	NM_002245		GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGT
MET-2	N/A	NM_002245		GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGT
MET-3	N/A	NM_002245		GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGT
MET-4	N/A	NM_002245		GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGT
MET-5	N/A	NM_002245		GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGT
MET-6	N/A	NM_002245		GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGT
MET-7	N/A	NM_002245		GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGT
MET-8	N/A	NM_002245		GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGT
MET-9	N/A	NM_002245		GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGT
MET-10	N/A	NM_002245		GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGT
MET-11	N/A	NM_002245		GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGT
MET-12	N/A	NM_002245		GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGT
MET-13	N/A	NM_002245		GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGT
MET-14	N/A	NM_002245		GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGT
MET-15	N/A	NM_002245		GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGT
MET-16	N/A	NM_002245		GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGT
MET-17	N/A	NM_002245		GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGT
MET-18	N/A	NM_002245		GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGT
MET-19	N/A	NM_002245		GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGT
MET-20	N/A	NM_002245		GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGT
NTRK1-1	CCDS1161.1	NM_002529.1	AAAGCGCACTTGTGAGGATTC	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGT
NTRK1-2	CCDS1161.1	NM_002529.1	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGT
NTRK1-3	CCDS1161.1	NM_002529.1	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGT
NTRK1-4	CCDS1161.1	NM_002529.1	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGT
NTRK1-5	CCDS1161.1	NM_002529.1	CCTCCCTTACCTGTGAGC	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGT
NTRK1-6	CCDS1161.1	NM_002529.1	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGT
NTRK1-7	CCDS1161.1	NM_002529.1	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGT
NTRK1-8	CCDS1161.1	NM_002529.1	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGT
NTRK1-9	CCDS1161.1	NM_002529.1	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGT
NTRK1-10	CCDS1161.1	NM_002529.1	ATGAGTCCAGAGTGGCAGG	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGT
NTRK1-11	CCDS1161.1	NM_002529.1	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGT
NTRK1-12	CCDS1161.1	NM_002529.1	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGT
NTRK1-13	CCDS1161.1	NM_002529.1	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGT
NTRK1-14	CCDS1161.1	NM_002529.1	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGT
NTRK1-15	CCDS1161.1	NM_002529.1	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGT
NTRK1-16	CCDS1161.1	NM_002529.1	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGT
NTRK1-17	CCDS1161.1	NM_002529.1	CGAGTAGTGTGAGCTGCTT	GTAAAACGCGGCCAGTCTGCTCATGACACAG	GTAAAACGCGGCCAGT

PTK7-20	CCDS4884.1	NM_002821.3	GTAAAACGCGGCCAGTTGCAAGGTCCACAAAGTGGT	AGACCTCAGCATGCCCTGTG	GTAAAACGCGGCCAGT
ROR2-1	CCDS6691.1	NM_004560.2	GTAAAACGCGGCCAGTGGACGCATCGTAGAAAGGG	GCTTATTGTAAACCGGCCAG	GTAAAACGCGGCCAGT
ROR2-2	CCDS6691.1	NM_004560.2	GTAAAACGACGGCCAGTCGGCATTGGCTATTCTTG	AAACACAGAATCAAGGTGGC	GTAAAACGCGGCCAGT
ROR2-3	CCDS6691.1	NM_004560.2	GTAAAACGACGGCCAGTCCTCTGGAAAGATCATAGC	TGCTGACTGGTGTGTTAG	GTAAAACGCGGCCAGT
ROR2-4	CCDS6691.1	NM_004560.2	GTAAAACGACGGCCAGTCCTCCCTTGTGTGGTTTC	CAAATTCCGAAAGACATGAG	GTAAAACGCGGCCAGT
ROR2-5	CCDS6691.1	NM_004560.2	AATTGCTGGATCGCAAGATG	GTAAAACGACGGCCAGTCCCACATGCCATTAAAC	GTAAAACGCGGCCAGT
ROR2-6	CCDS6691.1	NM_004560.2	GTAAAACGACGGCCAGTGTGGCAGTCTGGGATGC	CCTGGGCTTCACCGACAC	GTAAAACGACGGCCAGT
ROR2-7	CCDS6691.1	NM_004560.2	GTAAAACGACGGCCAGTATTGGGCTTGGACTTC	GTCAGGACAGAACGCCCT	GTAAAACGACGGCCAGT
ROR2-8	CCDS6691.1	NM_004560.2	GTAAAACGACGGCCAGTAGGTGGAGAGTGGGGTTAG	GCAGTGAATCCCCAACAG	GTAAAACGACGGCCAGT
ROR2-9	CCDS6691.1	NM_004560.2	GTAAAACGACGGCCAGTAGGAAGTCAGTGTGCCAGC	AGCTGAGAGATCATGCTCAGG	GTAAAACGACGCCAGT
ROR2-9	CCDS6691.1	NM_004560.2	GACCGTTTGGAAAGTCATAC	GTAAAACGACGGCCAGTCACCGTGGAGGAGATGTC	GTAAAACGACGCCAGT
ROR2-9	CCDS6691.1	NM_004560.2	AGTACCTATCACGGCCACACC	GTAAAACGACGGCCAGTGTGACATTTGCTACTG	GTAAAACGACGCCAGT
ROR2-9	CCDS6691.1	NM_004560.2	GTAAAACGACGGCCAGTCAGTTCTCATGCAGTCAG	ATTCGACATTGGATCTGCA	GTAAAACGACGCCAGT
ROR2-9	CCDS6691.1	NM_004560.2	CCAGTGACCAATGTGAGCACAC	GTAAAACGACGGCCAGTCAGTGGACTGAGGTCCC	GTAAAACGACGCCAGT
TIE1-1	CCDS482.1	NM_005424.2	GTAAAACGACGGCCAGTAGGCCACCTCATTCCTCTCC	AATTGATCTCCCATCTCAGC	GTAAAACGACGCCAGT
TIE1-2	CCDS482.1	NM_005424.2	GTAAAACGACGGCCAGTAGAAGACAGAGGGCTGG	GACCTGTTCTGAAGCAGGTG	GTAAAACGACGCCAGT
TIE1-3	CCDS482.1	NM_005424.2	ACAGCCCTGAGGTGAAGTTAG	GTAAAACGACGGCCAGTCACAGAGAGGCCCTGG	GTAAAACGACGCCAGT
TIE1-4	CCDS482.1	NM_005424.2	AGCTGAGCACGGGTGGACAG	GTAAAACGACGGCCAGTCAGGAAAGGTGCTTCAGC	GTAAAACGACGCCAGT
TIE1-5	CCDS482.1	NM_005424.2	GTAAAACGACGGCCAGTCGCCCTATGGGTACTCTGTG	CACCCCATGTTCTAGGGTC	GTAAAACGACGCCAGT
TIE1-6	CCDS482.1	NM_005424.2	GACCTAGAACCATGTGGT	GTAAAACGACGGCCAGTCAGTGGCAGAC	GTAAAACGACGCCAGT
TIE1-7	CCDS482.1	NM_005424.2	GTAAAACGACGGCCAGTCGGGATCTTCACCTCTCAGCC	AGAGCTCTCTGTCATCAGAC	GTAAAACGACGCCAGT
TIE1-8	CCDS482.1	NM_005424.2	GTAAAACGACGGCCAGTCACAGGAAACAGAACACG	CTGACCTATGTAAGTGG	GTAAAACGACGCCAGT
TIE1-9	CCDS482.1	NM_005424.2	GTAAAACGACGGCCAGTATCAGTCCAGCAGACCTG	AGTATCTCAGCCTGAGCAGC	GTAAAACGACGCCAGT
TIE1-10	CCDS482.1	NM_005424.2	GTCAAGCTTGAACAAAGATGC	GTAAAACGACGGCCAGTCAGTGGCTCC	GTAAAACGACGCCAGT
TIE1-11	CCDS482.1	NM_005424.2	GTAAAACGACGGCCAGTGTGAGTAGGGAGAGCTGGG	CAAGGAAGAACACCTCCAGTG	GTAAAACGACGCCAGT
TIE1-12	CCDS482.1	NM_005424.2	GTAAAACGACGGCCAGTTGTTCTCTGTGTTACTGGG	CCTCCCTGCTCTCTAGGTC	GTAAAACGACGCCAGT
TIE1-13	CCDS482.1	NM_005424.2	GTAAAACGACGGCCAGTTAGAGCAGATGGTCAGCC	TGTGGAAGGAGAGGGTGC	GTAAAACGACGCCAGT
TIE1-14	CCDS482.1	NM_005424.2	GTAAAACGACGGCCAGTGGGATCTTCACCTCTCC	CACACACAGGTGACAAGGTATTG	GTAAAACGACGCCAGT
TIE1-15	CCDS482.1	NM_005424.2	GTAAAACGACGGCCAGTGGGATCTTCACCTCTCC	GTCATTGCTGGAGGAAGCC	GTAAAACGACGCCAGT
TIE1-16	CCDS482.1	NM_005424.2	GTAAAACGACGGCCAGTTGGTCCACAAATCCAG	GATGAAAGGAGGTAAGGGT	GTAAAACGACGCCAGT
TIE1-17	CCDS482.1	NM_005424.2	GTAAAACGACGGCCAGTATCCCTGTCGTACCATCGG	GGCTGACTCTTCACTGACAC	GTAAAACGACGCCAGT
TIE1-18	CCDS482.1	NM_005424.2	CGAAAGACTGACTCTTACTGGC	GTAAAACGACGGCCAGTCCCTAGGGAGAGATGGGTTG	GTAAAACGACGCCAGT
TIE1-19	CCDS482.1	NM_005424.2	AGTCATTCATACCCCTCACC	GTAAAACGACGGCCAGTCACCTGTGTCACACTGGAAC	GTAAAACGACGCCAGT
TIE1-20	CCDS482.1	NM_005424.2	GTAAAACGACGGCCAGTTGCCAAGGCCAGACTTAC	ACACTTATGTCACAGTTG	GTAAAACGACGCCAGT
TIE1-21	CCDS482.1	NM_005424.2	GTAAAACGACGGCCAGTGGGAGATAGTGAGCCTTGTG	TAGCTGTAGCGCAATCTGG	GTAAAACGACGCCAGT
TIE1-22	CCDS482.1	NM_005424.2	GTAAAACGACGGCCAGTCAGCTTGTGACCTCTGTGACC	GCCCTGGTGTAAACCAAGTGG	GTAAAACGACGCCAGT
TIE1-23	CCDS482.1	NM_005424.2	GTAAAACGACGGCCAGTAGGAAGTCCAGGAGCTTGAG	CACCTACAAAGCATGGGAAAC	GTAAAACGACGCCAGT

#The primer pair did not meet our quality criteria that ≥90% of bases in the target region have a Phred quality score of at least 20 in three quarters of the tumor samples analyzed.

Table S4. Characteristics of melanoma patients with PTK mutations

Sample	Patient Age (years)*	Patient Gender	Tumor Source	Matched normal source
1T	29	F	Lung	Blood
2T	30	M	Pectoral muscle	Blood
3T	18	M	Forehead, subcutaneous	Blood
4T	33	F	Supraclavicular, soft tissue	Blood
4T	34	F	Lung	Blood
5T	47	M	External iliac soft tissue	Blood
6T	42	M	Neck, soft tissue	Blood
7T	53	M	Stomach	Blood
8T	61	M	Inguinal soft tissue	Blood
9T	62	M	Back, subcutaneous	Blood
10T	55	M	Axillary soft tissue	Blood
12T	53	M	Upper arm, subcutaneous	Blood
13T	49	M	Chest wall, subcutaneous	Blood
15T	39	M	Thigh, subcutaneous	Blood
16T	62	M	Lung	Blood
17T	33	M	Shoulder, subcutaneous	Blood
18T	55	M	Clavicle, soft tissue	Blood
19T	49	M	Scapula, subcutaneous	Blood
20T	58	F	Axillary soft tissue	Blood
21T	59	M	Omentum	Blood
22T	51	M	Chest wall, subcutaneous	Blood
23T	44	M	Lung	Blood
24T	49	M	Axillary soft tissue	Blood
26T	48	F	Lung	Blood
28T	28	F	Axillary soft tissue	Blood
29T	51	M	Inguinal soft tissue	Blood
30T	53	F	Lung	Blood
31T	49	F	Thigh, subcutaneous	Blood
32T	58	M	Omentum	Blood
33T	33	M	Chest wall subcutaneous, & pleura **	Blood
34T	31	M	Shoulder, subcutaneous	Blood
35T	23	F	Thigh, subcutaneous	Blood
36T	25	M	Thigh, subcutaneous	Blood
37T	38	F	Omentum	Blood
38T	27	M	Skull	Blood
39T	56	M	Mesentery	Blood
41T	45	M	Neck, soft tissue	Blood
43T	19	F	Popliteal soft tissue	Blood
44T	56	M	Lung	Blood
45T	48	M	Mediastinum	Blood
49T	43	M	Thigh, subcutaneous	Blood
50T	49	F	Inguinal soft tissue	Blood
51T	50	F	Adnexa	Blood
52T	39	F	Lung	Blood
54T	27	M	Subcutaneous	Blood
55T	60	M	Lung	Blood
56T	52	M	Lung	Blood
58T	46	F	Hip, subcutaneous	Blood
59T	64	F	Abdomen, subcutaneous	Blood
60T	46	M	Flank, subcutaneous	Blood
62T	58	F	Thigh, subcutaneous	Blood
63T	30	M	Jejunum	Blood
64T	32	F	Ovary	Blood
68T	49	M	Lung	Blood
69T	36	M	Axillary soft tissue	Blood
71T	67	M	Lung	Blood
72T	53	M	Liver	Blood
73T	45	F	Breast	Blood
74T	40	F	Lower extremity, subcutaneous	Blood
76T	40	M	Neck, soft tissue	Blood
77T	39	M	Lung	Blood
78T	27	F	Lung	Blood
79T	53	M	Supraclavicular, soft tissue	Blood
80T	36	F	Popliteal soft tissue	Blood
81T	60	F	Upper arm, subcutaneous	Blood
83T	33	F	Back, subcutaneous	Blood
84T	60	F	Thigh, subcutaneous	Blood
85T	44	M	Chest wall, subcutaneous	Blood
86T	42	F	Liver	Blood
87T	27	M	Small bowel & mesentary **	Blood
88T	37	F	Chest wall, subcutaneous	Blood
90T	19	M	Neck, soft tissue	Blood
91T	55	F	Subcostal soft tissue	Blood
92T	37	F	Femur	Blood
93T	42	F	Axillary soft tissue	Blood
94T	44	M	Adrenal gland	Blood
95T	58	F	Inguinal soft tissue	Blood
96T	49	M	Inguinal soft tissue	Blood
99T	57	M	Liver	Blood

* Patient's age when tumor was surgically removed. ** Cell line generated by cells mixed from 2 tumors resected during the patient's surgery. Abbreviations: F, female; M, male.

Table S5. Mutations Identified in RAF and RAS isoforms in melanoma samples containing ERBB4 mutations

Sample	ERBB4	BRAF	NRAS	ARAF	CRAF	HRAS	KRAS
7T	E452K	wt	Q61R	wt	wt	wt	wt
12T	E563K	wt	Q61Q/R	wt	wt	wt	wt
17T	E317K	wt	Q61Q/K	wt	wt	wt	wt
31T	wt	wt	wt	wt	wt	wt	wt
34T	R491K	V600V/E	wt	wt	T362T/A	wt	wt
39T	wt	wt	wt	wt	wt	wt	wt
49T	R393R/W	V600V/E	wt	wt	wt	wt	wt
55T	E452K	V600V/E	wt	P216S, P254L	wt	wt	wt
56T	R544R/W	V600V/E	wt	wt	wt	wt	wt
63T	E542K, E872K	wt	Q61Q/K	wt	wt	wt	wt
68T	Splice Site / LOH	V600V/E	wt	wt	wt	wt	wt
71T	L39L/F, S1246S/N	V600V/M, V600V/E	wt	wt	wt	wt	wt
86T	E836E/K	V600V/E	wt	wt	wt	wt	wt
93T	wt	wt	wt	A345A/G	wt	wt	wt

Table S6. Primers used for PCR amplification and sequencing of RAS and Raf isoforms

Gene and Exon Name	Forward Primer	Reverse Primer	Sequencing Primer
ARAF-1	TGAGCAGGGATCTTGGACTG	GTAaaaACGAGGCCAGTCAGCGATGTTCCCACCTTC	GTAAAACGACGCCAGT
ARAF-2	GTCATGGAACTGG	GTAAAACGACGCCAGTCAGAGGATCACTGACTGAGG	GTAAAACGACGCCAGT
ARAF-3	GTAAAAACGACGCCAGTTACCTCTGAGCCCTGTTCC	ACGAGGGAGTTTGAGAACCTG	GTAAAACGACGCCAGT
ARAF-4	GTAAAAACGACGCCAGTACCAACCTCCACTCATTC	CACGGGTGAGCTGTTGAAG	GTAAAACGACGCCAGT
ARAF-5	AGTACCAACCCCAAACAGT	GTAAAACGACGCCAGTGGAAAATGAGGTGACTCTGCC	GTAAAACGACGCCAGT
ARAF-6	CAATTCATGTTTATGGCTGG	GTAAAACGACGCCAGTCAGTGTGAAACTCTGGC	GTAAAACGACGCCAGT
ARAF-7	GTAAAACGACGCCAGTCACGCTCATATGGTCAGCAC	CTGTTGACTTGGAAATGGGG	GTAAAACGACGCCAGT
ARAF-8	CCAGAGTTCAGCAATGGT	GTAAAACGACGCCAGTCAGCTGAGCTGAGG	GTAAAACGACGCCAGT
ARAF-9	GTAAAACGACGCCAGTAAAGTGATGCTCGAGGG	ATGTCAGGAACTCACTGG	GTAAAACGACGCCAGT
ARAF-10	GTAAAACGACGCCAGTACAGCTGGACTCTGTATG	TTCTGGTATGGATTTCTTG	GTAAAACGACGCCAGT
ARAF-11	GGATTGTCATCATCACAG	GTAAAACGACGCCAGTCTTGTGATTCATCCCCAGACCC	GTAAAACGACGCCAGT
ARAF-12	GTAAAACGACGCCAGTGGATGTTGAGTAGTTGCG	GTGAGGATCACTGGAGCTTCAAG	GTAAAACGACGCCAGT
ARAF-13	GTGTTGGTGTGCTGTTG	GTAAAACGACGCCAGTGGTATTCATGCGAAAGG	GTAAAACGACGCCAGT
ARAF-14	GTAAAACGACGCCAGTACAGCAATCTCCAAAGTC	TCACATCTGCTCATCTCG	GTAAAACGACGCCAGT
ARAF15 ^a	CTCTGTTTACATCAAGACCC	GTAAAACGACGCCAGTCACACACCAATTGATTG	GTAAAACGACGCCAGT
BRaf-1	GTAAAACGACGCCAGTACGGCTCCCTCC	GTAAAACGACGCCAGTCACGGCATC	GTAAAACGACGCCAGT
BRaf-2	GTAAAACGACGCCAGTGGAAACACTGGCAGTTCTG	TCTCTCCCAATCTATTCTTAACTC	GTAAAACGACGCCAGT
BRaf-3	TGGTTGTTGATCTGACCTGTAACT	GTAAAACGACGCCAGTCCTCTCCTCTTC	GTAAAACGACGCCAGT
BRaf-4	GTAAAACGACGCCAGTCCCTACTGTTACTGAGCC	TTACATCATCATTCATCTCC	GTAAAACGACGCCAGT
BRaf-5	GTAAAACGACGCCAGCTGTAGTCTATTATGGAA	GGAGAAGGATTCCTTCTTGC	GTAAAACGACGCCAGT
BRaf-6	GTGTTTGTGAGATGGATTG	GTAAAACGACGCCAGTTCTGTTGATGATGTTTGGG	GTAAAACGACGCCAGT
BRaf-7	GTAAAACGACGCCAGTACGGCTGGAGATTGGATT	TOCATAGAGGAGGAGGAGG	GTAAAACGACGCCAGT
BRaf-8	GTAAAACGACGCCAGTGGTACAGGTTGG	GTAACTTGAAGAGGAGGAGTAA	GTAAAACGACGCCAGT
BRaf-9	GTAAAACGACGCCAGTGGTACACATGGCTTC	GGTCTGGCGTAAATAATGTTG	GTAAAACGACGCCAGT
BRaf-10	GTAAAACGACGCCAGTGGTACCTCC	GTAAAACGACGCCAGTCACAGTGTGATG	GTAAAACGACGCCAGT
BRaf-11	GTAAAACGACGCCAGTCATGGAAACAAAGGTTG	AATAGTTGCAACCTGGAAAC	GTAAAACGACGCCAGT
BRaf-12	CAATTCAGCTCATCTCTAATGATC	GTAAAACGACGCCAGTTGATCCTCTATGCTTGGAC	GTAAAACGACGCCAGT
BRaf-13	GTAAAACGACGCCAGTCATGGAAATATGGCTACTG	CCATACATGCGACAACTCC	GTAAAACGACGCCAGT
BRaf-14	TCTCATGACACATTTCAGGCC	GTAAAACGACGCCAGTTGTAATGAGGAACTTAA	GTAAAACGACGCCAGT
BRaf-15	GAATTCATCTGGCATGATGTT	GTAAAACGACGCCAGTGGCAACCCAGGAA	GTAAAACGACGCCAGT
BRaf-16	CCATCTGATGCGATGTT	GTAAAACGACGCCAGTGTGAAACCCATC	GTAAAACGACGCCAGT
BRaf-17	GCTTTCTGTAAAGTGTGATG	GTAAAACGACGCCAGTCCACAAAGCTGTTCTGGTC	GTAAAACGACGCCAGT
RAF1-1	GTAAAACGACGCCAGTGTGCTTACCTTCTCATC	GTAAAACGACGCCAGTGGGAAAC	GTAAAACGACGCCAGT
RAF1-2	GTAAAACGACGCCAGTATTCTGTCGACCCCTTC	AGGTATTGGTCAGGGCCC	GTAAAACGACGCCAGT
RAF1-3	GTAAAACGACGCCAGTGGCTTGGAAACAACTTC	TTGCCCTACTGTAACACAGCA	GTAAAACGACGCCAGT
RAF1-4	GTAAAACGACGCCAGTGGGCAAGAATGAGGTT	ATGAATGCCACCAACTTCG	GTAAAACGACGCCAGT
RAF1-5	GTAAAACGACGCCAGTGGGCAAGTCAGTGTGTT	CCACAGAAAGCAGCAAGG	GTAAAACGACGCCAGT
RAF1-6	GTAAAACGACGCCAGTGTGATGAGGTTGGCC	TCCTCTGATCATTTGAAACC	GTAAAACGACGCCAGT
RAF1-7	GTAAAACGACGCCAGTGTGAGGTTGGAGA	TTGGCAGAGGAGTGTGTC	GTAAAACGACGCCAGT
RAF1-8	GGATCCATTGCAAGTCACAG	GTAAAACGACGCCAGTCTCTCCCTCT	GTAAAACGACGCCAGT
RAF1-9	AAACAGCATGGTGTGATCC	GGCGCACGTCACAACT	GTAAAACGACGCCAGT
RAF1-10	GTAAAACGACGCCAGTTGGAAATTGCGCTATG	GGTTGTGCAAGAATTCACAG	GTAAAACGACGCCAGT
RAF1-11	GTAAAACGACGCCAGTGGCAACGACCCACTTC	TTCTCTGTCCTCTGCTTTC	GTAAAACGACGCCAGT
RAF1-12	GTAAAACGACGCCAGTCGCTCTGTGTAACACTCTGG	GTAAAACGACGCCAGTAACTCAAAATTGCTGAGG	GTAAAACGACGCCAGT
RAF1-13	GCTGTGACAGGTTAAAGTGTG	AGGCTCTTCAATTGTTTGGG	GTAAAACGACGCCAGT
RAF1-14	GTAAAACGACGCCAGTCATAAGGAAAGCAGCTGG	GTAAAACGACGCCAGTCCATTGTTGGG	GTAAAACGACGCCAGT
RAF1-15	CAGTGAATCTGTCGTTG	GTAAAACGACGCCAGTCCATTGAGGACCTGG	GTAAAACGACGCCAGT
RAF1-16	CCACGATCTCTCAAGATG	GTAAAACGACGCCAGTGTGTTCTGCTCTGG	GTAAAACGACGCCAGT
HRAS-1	TGGGTCTTAAAGGATGTTG	GTAAAACGACGCCAGTGTGAGGACGAGACAG	GTAAAACGACGCCAGT
HRAS-2	GTAAAACGACGCCAGTGGAGCTGGCTGTGTGAAC	GACATGGCGAGAGAGGAC	GTAAAACGACGCCAGT
HRAS-3	TTCTCTGTTCTTCATC	GTAAAACGACGCCAGTGGTGTGCTCTCTGG	GTAAAACGACGCCAGT
HRAS-4	GGCTCTGTCCTCTC	GTAAAACGACGCCAGTGTGCTCTCCAAAGGACCT	GTAAAACGACGCCAGT
KRAS-1	GTAAAACGACGCCAGTGGAGCTGAACGAGTGGT	AGAAAGCAGCTGAAGGAGGAG	GTAAAACGACGCCAGT
KRAS-2	GTAAAACGACGCCAGTCGTCGGATGAGGATAGG	TTTCATGCTCTCTTCC	GTAAAACGACGCCAGT
KRAS-3	GTAAAACGACGCCAGTGGAGCTGGCATAGTGGCTGAC	ACTCGAGTACAGCACGCC	GTAAAACGACGCCAGT
KRAS-4	ATTTCCACATTGCGGCTGAG	GTAAAACGACGCCAGTGGCAATGTTGGAAAG	GTAAAACGACGCCAGT
NRAS-1	GTAAAACGACGCCAGTAAATGCGATTGCTTCCCTG	GAACATACAGTGGTCAATAG	GTAAAACGACGCCAGT
NRAS-2	GTAAAACGACGCCAGTAAATGCGATTGCTTCCCTG	CTCTGTTCCAAGCTTCC	GTAAAACGACGCCAGT
NRAS-3	GTAAAACGACGCCAGTAAATGCGATTGCTTCCCTG	CAAGAGACAGAGGCTCAGT	GTAAAACGACGCCAGT
NRAS-4	GCGCTGTTCTGTGATTCATAG	GTAAAACGACGCCAGTGGCAAGGAGGATAGGCG	GTAAAACGACGCCAGT

^aThe primer pair did not meet our quality criteria that ≥90% of bases in the target region have a Phred quality score of at least 20 in three quarters of the tumor samples analyzed.

Table S7. Primers used for ERBB4 plasmid construction

Gene Name	Forward Primer	Reverse Primer
ERBB4 into pCDF-MCS2-EF1-Puro™	CGGCTCTAGAGGCCACCATGAGGCCGGCAG	ATCGGCCGGCGCTTACCCACAGTATCCGG
Knockdown resistant ERBB4 primers into pCDF1-MCS2-EF1-puro	CACCAAAATCAAGGGGACTT	GGAGCCAGTACACGACATCA
Knockdown resistant ERBB4 primer set #1 into pcDNA3.1	GATTCCTGTCGGCTTAAAGATTCT	AGAACTTAATGGCCACAGGAATC
Knockdown resistant ERBB4 primer set #2 into pcDNA3.1	CCTCTGGCCATCAAGATTCT	AGATCTTGTGATGGCCACAGG
Knockdown resistant ERBB4 primer set #3 into pcDNA3.1	CCTCTGGCCATCAAAATTCTTAATGAGAC	GTCTCATTAAGAATTGATGGCCACAGG

Supplementary Note**Tumor Tissue Collection**

A panel of 79 pathology-confirmed metastatic melanoma tumor resections was assembled from patients referred to the Surgery Branch of the National Cancer Institute for enrollment in IRB-approved Surgery Branch clinical trials of immunotherapy. All of the patients had progressive metastatic melanoma that had failed available conventional treatments, were 16 years or older and passed eligibility testing for the relevant protocol. None had received therapy for 1 month prior to entering the protocol. Before tissue was acquired all patients signed written informed consents. The melanoma tissue panel was selected on the basis of an available pathology-confirmed, melanoma tissue culture line, paired with freshly frozen tissue from the resected melanoma metastasis and apheresis-collected, peripheral blood mononuclear cells.