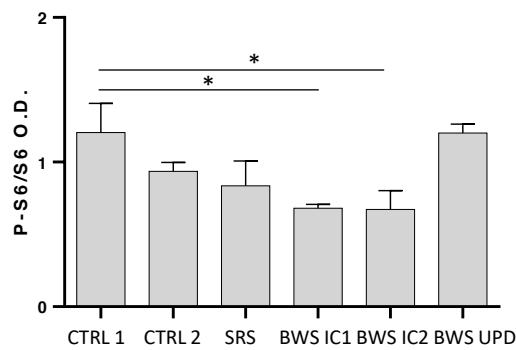
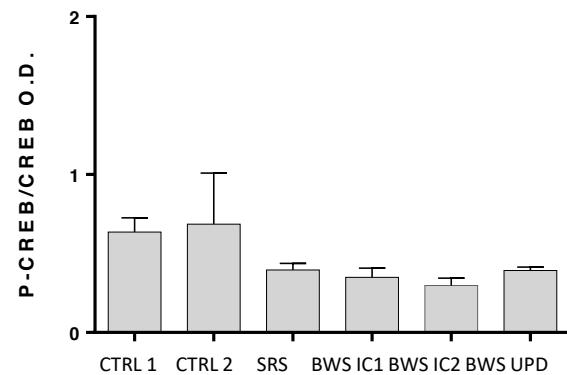
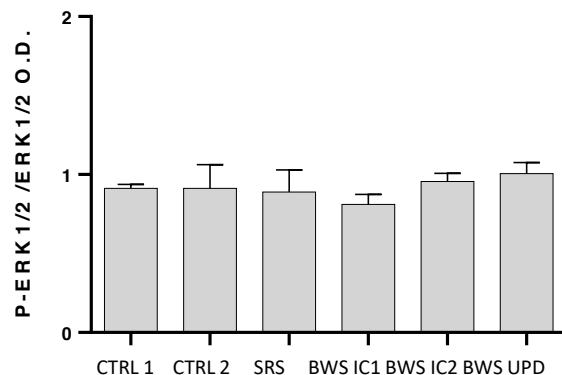
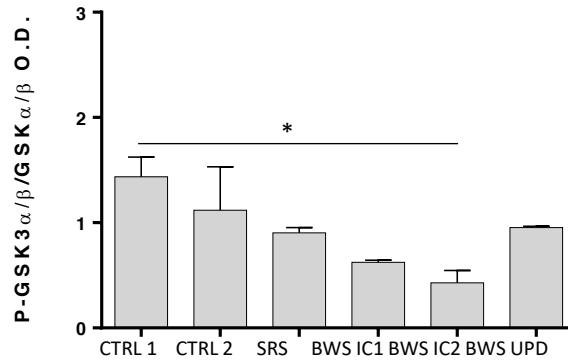
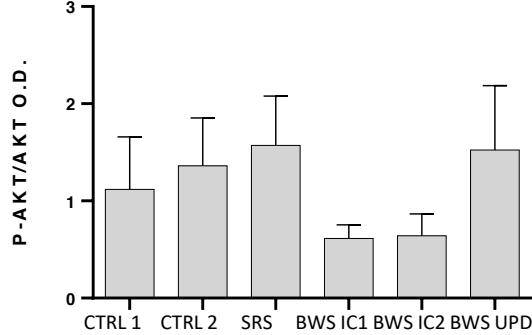


**Supplemental Figure S1.** Analysis of the phosphorylation pattern of components of the PI3K and MAPK pathways by PathScan EGFR signaling antibody array Kit in BWS IC2 and CTRL LCLs. Phosphorylation of MEK1/2 (Ser221) and MEK1/2 (Ser217/221) are highlighted in blue and green, respectively. Phosphorylation of Akt (Thr308) and Akt (Ser473) are highlighted in yellow and orange, respectively.



**Supplemental Figure S2.** Relative intensity (Optical density: O.D.) by densitometric analysis was evaluated and expressed as ratio of phospho/total proteins. Data represent the mean  $\pm$  SEM of 3 experiments (two way-ANOVA, \* $<0.05$ )

Table S1. Panel of genes of the WNT pathway analyzed by nCounter approach

GENE	Accession	Log2 fold change	P-value	BH.p.value	Pathway annotation
<i>TP53</i>	NM_000546.2	0.632	2.81e-05	0.00446	KEGG WNT Annotation
<i>PLAUR</i>	NM_001005376.1	-2.16	0.000288	0.0229	Proteolysis
<i>CUL1</i>	NM_003592.2	0.196	0.000798	0.0377	KEGG WNT Annotation
<i>PRKCB</i>	NM_212535.1	-1.23	0.00102	0.0377	KEGG WNT Annotation
<i>FRAT1</i>	NM_005479.3	1.83	0.00146	0.0377	Canonical Wnt Pathway, KEGG WNT Annotation
<i>AXIN1</i>	NM_181050.1	0.265	0.00149	0.0377	Canonical Wnt Pathway, KEGG WNT Annotation, WNT Signaling Negative Regulation
<i>MAPK9</i>	NM_139068.2	0.51	0.00166	0.0377	KEGG WNT Annotation
<i>CEBPD</i>	NM_005195.3	1.58	0.0027	0.0481	Transcription Factors
<i>DKK4</i>	NM_014420.2	-1.2	0.00272	0.0481	KEGG WNT Annotation
<i>MAPK10</i>	NM_002753.2	-6.57	0.00526	0.079	KEGG WNT Annotation
<i>CREBBP</i>	NM_001079846.1	0.547	0.00573	0.079	KEGG WNT Annotation
<i>PITX2</i>	NM_000325.5	3.94	0.00649	0.079	Transcription Factors, WNT Signaling Target Genes
<i>RBX1</i>	NM_014248.2	-0.521	0.00682	0.079	KEGG WNT Annotation
<i>MMP9</i>	NM_004994.2	2.09	0.00722	0.079	Calcium Binding and Signaling, Development & Differentiation, Migration, Proteolysis
<i>FBXW11</i>	NM_033645.2	0.313	0.00745	0.079	KEGG WNT Annotation, WNT Signaling Negative Regulation
<i>WNT10A</i>	NM_025216.2	-2.25	0.00837	0.0832	Calcium Binding and Signaling, Canonical Wnt Pathway, KEGG WNT Annotation
<i>GSK3A</i>	NM_019884.2	0.167	0.0168	0.157	Canonical Wnt Pathway

<i>GDNF</i>	NM_000514.2	-4.12	0.0222	0.185	Development & Differentiation, Migration
<i>SMAD2</i>	NM_005901.5	0.278	0.0268	0.206	EMTMetastasis, KEGG WNT Annotation
<i>SMAD4</i>	NM_005359.3	0.374	0.0272	0.206	KEGG WNT Annotation
<i>ERBB2</i>	NM_001005862.1	-0.679	0.0319	0.222	EMTMetastasis
<i>PKN1</i>	NM_213560.1	0.357	0.0322	0.222	EMTMetastasis
<i>PTGS2</i>	NM_000963.1	-2.07	0.0335	0.222	Calcium Binding and Signaling, Cell Cycle
<i>CXCL12</i>	NM_000609.5	-3.36	0.0367	0.226	EMTMetastasis
<i>BMP4</i>	NM_001202.3	0.757	0.0372	0.226	Development & Differentiation
<i>T</i>	NM_003181.2	4.22	0.0383	0.226	Development & Differentiation, Transcription Factors
<i>BAMBI</i>	NM_012342.2	3.2	0.0408	0.226	Canonical Wnt Pathway
<i>CCND1</i>	NM_053056.2	-2.39	0.0425	0.226	Calcium Binding and Signaling, Cell Cycle, Development & Differentiation, KEGG WNT Annotation, WNT Signaling Negative Regulation, WNT Signaling Target Genes
<i>FZD2</i>	NM_001466.2	-2.36	0.0426	0.226	Calcium Binding and Signaling, Canonical Wnt Pathway, KEGG WNT Annotation
<i>PRKACA</i>	NM_002730.3	-0.319	0.0519	0.252	KEGG WNT Annotation
<i>FOXN1</i>	NM_003593.2	-3.47	0.055	0.252	Development & Differentiation
<i>TCF7L1</i>	NM_031283.1	-1.11	0.0575	0.252	Canonical Wnt Pathway, KEGG WNT Annotation, Transcription Factors
<i>TBL1XR1</i>	NM_024665.4	0.332	0.061	0.252	KEGG WNT Annotation
<i>ETS2</i>	NM_005239.4	0.771	0.0618	0.252	Transcription Factors
<i>WNT10B</i>	NM_003394.2	-1.67	0.0627	0.252	KEGG WNT Annotation
<i>MYC</i>	NM_002467.3	0.568	0.0639	0.252	Cell Cycle, KEGG WNT Annotation, Transcription Factors, WNT Signaling Target Genes

<i>WNT2</i>	NM_003391.2	-1.47	0.0647	0.252	Calcium Binding and Signaling, Canonical Wnt Pathway, KEGG WNT Annotation
<i>FZD1</i>	NM_003505.1	1.58	0.0668	0.252	Canonical Wnt Pathway, KEGG WNT Annotation
<i>ZEB1</i>	NM_001128128.1	0.351	0.0673	0.252	EMTMetastasis
<i>SOX2</i>	NM_003106.2	-2.25	0.0691	0.252	Cell Cycle, Development & Differentiation, Transcription Factors
<i>POU5F1</i>	NM_002701.4	-0.589	0.0704	0.252	Development & Differentiation, Transcription Factors
<i>FN1</i>	NM_212482.1	-2.03	0.0706	0.252	Development & Differentiation, Migration
<i>FZD3</i>	NM_017412.2	-1.04	0.0713	0.252	Canonical Wnt Pathway, KEGG WNT Annotation
<i>FZD8</i>	NM_031866.1	-3.06	0.0742	0.256	Canonical Wnt Pathway, KEGG WNT Annotation
<i>TIMP1</i>	NM_003254.2	-1.19	0.0766	0.259	EMTMetastasis
<i>SERPINE1</i>	NM_001165413.1	-2.58	0.0809	0.268	EMTMetastasis
<i>PDGFRA</i>	NM_006206.3	-1.69	0.0855	0.278	Development & Differentiation, Migration
<i>APC</i>	NM_000038.3	0.293	0.0896	0.28	Canonical Wnt Pathway, KEGG WNT Annotation, WNT Signaling Negative Regulation
<i>RUVBL1</i>	NM_003707.2	-0.333	0.0908	0.28	Canonical Wnt Pathway, KEGG WNT Annotation
<i>RUNX2</i>	NM_004348.3	-0.691	0.0945	0.284	Development & Differentiation, Transcription Factors
<i>TCF4</i>	NM_003199.1	0.362	0.105	0.309	Transcription Factors
<i>LEF1</i>	NM_016269.3	1.58	0.11	0.316	Canonical Wnt Pathway, KEGG WNT Annotation, Transcription Factors
<i>CSNK1A1</i>	NM_001892.4	0.219	0.111	0.316	Canonical Wnt Pathway, KEGG WNT Annotation
<i>SOX9</i>	NM_000346.2	0.811	0.119	0.326	Adhesion, Development & Differentiation, Transcription Factors
<i>DVL1</i>	NM_004421.2	0.306	0.127	0.339	Canonical Wnt Pathway, KEGG WNT Annotation, Planar Cell Polarity (PCP)
<i>SMAD3</i>	NM_005902.3	0.887	0.132	0.34	KEGG WNT Annotation

<i>NRP1</i>	NM_003873.5	-4.33	0.134	0.34	Development & Differentiation, Migration
<i>PORCN</i>	NM_022825.2	0.22	0.135	0.34	Canonical Wnt Pathway, KEGG WNT Annotation
<i>DVL2</i>	NM_004422.2	0.229	0.137	0.34	Canonical Wnt Pathway, KEGG WNT Annotation, Planar Cell Polarity (PCP)
<i>AXIN2</i>	NM_004655.3	2	0.14	0.343	Canonical Wnt Pathway, KEGG WNT Annotation, WNT Signaling Negative Regulation, WNT Signaling Target Genes
<i>PRKACB</i>	NM_182948.2	0.836	0.143	0.345	KEGG WNT Annotation
<i>SNAI2</i>	NM_003068.3	-2.53	0.145	0.345	EMTMetastasis
<i>PPP3CB</i>	NM_001142354.1	-0.129	0.15	0.351	KEGG WNT Annotation
<i>KREMEN1</i>	NM_001039570.1	-1.67	0.157	0.351	WNT Signaling Negative Regulation
<i>SFRP1</i>	NM_003012.3	-1.58	0.159	0.351	Canonical Wnt Pathway, KEGG WNT Annotation, WNT Signaling Negative Regulation
<i>CTNNB1</i>	NM_001098210.1	0.221	0.159	0.351	Canonical Wnt Pathway, KEGG WNT Annotation
<i>DAB2</i>	NM_001343.2	-2.05	0.16	0.351	Development & Differentiation, WNT Signaling Target Genes
<i>WNT11</i>	NM_004626.2	2.15	0.163	0.351	Calcium Binding and Signaling, KEGG WNT Annotation
<i>NLK</i>	NM_016231.2	0.267	0.164	0.351	KEGG WNT Annotation, WNT Signaling Negative Regulation
<i>RAC1</i>	NM_198829.1	0.213	0.168	0.351	KEGG WNT Annotation
<i>CDKN2A</i>	NM_000077.3	0.552	0.168	0.351	Cell Cycle, Development & Differentiation, Transcription Factors
<i>CXXC4</i>	NM_025212.1	-0.783	0.172	0.355	Canonical Wnt Pathway, KEGG WNT Annotation, WNT Signaling Negative Regulation
<i>KLF5</i>	NM_001730.3	1.01	0.195	0.392	Transcription Factors
<i>PRKCG</i>	NM_002739.3	-0.992	0.198	0.394	KEGG WNT Annotation
<i>NANOG</i>	NM_024865.2	-0.367	0.206	0.404	Development & Differentiation, Transcription Factors
<i>IRS1</i>	NM_005544.2	0.815	0.212	0.41	Migration

<i>SKP1</i>	NM_170679.2	-0.09	0.217	0.413	KEGG WNT Annotation
<i>PPP3CA</i>	NM_000944.4	0.758	0.218	0.413	KEGG WNT Annotation
<i>RAC3</i>	NM_005052.2	0.337	0.221	0.413	KEGG WNT Annotation
<i>WNT4</i>	NM_030761.3	-1.03	0.224	0.414	Calcium Binding and Signaling, Canonical Wnt Pathway, KEGG WNT Annotation
<i>JUN</i>	NM_002228.3	0.501	0.233	0.421	KEGG WNT Annotation, WNT Signaling Target Genes
<i>TGFB3</i>	NM_003239.2	0.398	0.254	0.455	Development & Differentiation
<i>CCND3</i>	NM_001760.2	0.342	0.279	0.493	KEGG WNT Annotation
<i>MMP7</i>	NM_002423.3	-1.71	0.293	0.502	Calcium Binding and Signaling, KEGG WNT Annotation, Proteolysis, WNT Signaling Target Genes
<i>SIX1</i>	NM_005982.3	-1.61	0.294	0.502	Development & Differentiation, Migration, Transcription Factors
<i>PPARD</i>	NM_006238.3	-0.146	0.299	0.506	Development & Differentiation, KEGG WNT Annotation, Migration, Transcription Factors, WNT Signaling Target Genes
<i>FZD6</i>	NM_003506.2	0.818	0.315	0.527	Canonical Wnt Pathway, KEGG WNT Annotation
<i>IL6</i>	NM_000600.1	-1.1	0.328	0.543	Development & Differentiation, Migration
<i>VEGFA</i>	NM_001025366.1	0.954	0.355	0.578	Development & Differentiation, Migration
<i>AES</i>	NM_001130.5	0.126	0.357	0.578	Canonical Wnt Pathway
<i>WNT5A</i>	NM_003392.3	-1	0.368	0.586	Calcium Binding and Signaling, KEGG WNT Annotation
<i>WNT6</i>	NM_006522.3	1.23	0.368	0.586	Calcium Binding and Signaling, Canonical Wnt Pathway, KEGG WNT Annotation
<i>BIRC5</i>	NM_001168.2	-0.393	0.376	0.587	Cell Cycle
<i>CDH11</i>	NM_001797.2	-1.44	0.377	0.587	EMTMetastasis

<i>WNT7B</i>	NM_058238.1	0.836	0.394	0.609	Calcium Binding and Signaling, Canonical Wnt Pathway, KEGG WNT Annotation
<i>NRCAM</i>	NM_005010.4	-0.83	0.405	0.62	Adhesion, Development & Differentiation, Migration
<i>NKD1</i>	NM_033119.3	-0.322	0.419	0.635	Canonical Wnt Pathway, KEGG WNT Annotation, Planar Cell Polarity (PCP), WNT Signaling Negative Regulation
<i>FBXW4</i>	NM_022039.3	-0.154	0.449	0.667	WNT Signaling Negative Regulation
<i>CSNK2A1</i>	NM_177559.2	0.0641	0.453	0.667	Canonical Wnt Pathway, KEGG WNT Annotation
<i>FZD5</i>	NM_003468.2	0.582	0.47	0.683	Canonical Wnt Pathway, KEGG WNT Annotation
<i>LRP5</i>	NM_002335.1	0.302	0.472	0.683	Canonical Wnt Pathway, KEGG WNT Annotation
<i>FOSL1</i>	NM_005438.2	-0.404	0.501	0.712	Cell Cycle, KEGG WNT Annotation, Transcription Factors, WNT Signaling Target Genes
<i>SNAI1</i>	NM_005985.2	0.24	0.569	0.797	EMTMetastasis
<i>MYLK</i>	NM_053032.2	-0.175	0.572	0.797	EMTMetastasis
<i>CAMK2B</i>	NM_001220.3	0.369	0.588	0.81	KEGG WNT Annotation
<i>CTBP1</i>	NM_001328.2	-0.0632	0.591	0.81	Canonical Wnt Pathway, KEGG WNT Annotation, WNT Signaling Negative Regulation
<i>NFATC1</i>	NM_172389.1	-0.36	0.601	0.812	Calcium Binding and Signaling, KEGG WNT Annotation
<i>GPC4</i>	NM_001448.2	1.71	0.602	0.812	Canonical Wnt Pathway
<i>PRKCA</i>	NM_002737.2	0.471	0.626	0.826	KEGG WNT Annotation
<i>PLCB1</i>	NM_182734.1	-0.534	0.629	0.826	KEGG WNT Annotation
<i>DIXDC1</i>	NM_001037954.3	-0.176	0.639	0.833	Canonical Wnt Pathway
<i>EFNB1</i>	NM_004429.4	0.197	0.654	0.845	Development & Differentiation, Migration
<i>PPP3R1</i>	NM_000945.3	0.0488	0.667	0.848	KEGG WNT Annotation
<i>BCL9</i>	NM_004326.2	-0.132	0.681	0.857	Canonical Wnt Pathway

<i>EP300</i>	NM_001429.2	0.057	0.688	0.857	Canonical Wnt Pathway, KEGG WNT Annotation
<i>WNT2B</i>	NM_024494.1	-0.256	0.696	0.857	Calcium Binding and Signaling, Canonical Wnt Pathway, KEGG WNT Annotation
<i>MAPK8</i>	NM_002750.2	0.117	0.707	0.857	KEGG WNT Annotation, Planar Cell Polarity (PCP)
<i>JAG1</i>	NM_000214.2	0.266	0.709	0.857	Development & Differentiation, Migration
<i>ID2</i>	NM_002166.4	-0.276	0.718	0.857	Cell Cycle, Development & Differentiation
<i>PPP3CC</i>	NM_005605.4	0.0829	0.722	0.857	KEGG WNT Annotation
<i>SMO</i>	NM_005631.3	-0.28	0.732	0.857	Development & Differentiation, Migration
<i>GSK3B</i>	NM_002093.2	0.0354	0.735	0.857	Canonical Wnt Pathway, KEGG WNT Annotation
<i>ZEB2</i>	NM_014795.3	0.207	0.735	0.857	EMTMetastasis
<i>RHOA</i>	NM_001664.2	-0.0369	0.737	0.857	KEGG WNT Annotation, Planar Cell Polarity (PCP)
<i>CD44</i>	NM_001001392.1	0.0591	0.739	0.857	Adhesion, Migration
<i>FZD7</i>	NM_003507.1	0.33	0.753	0.86	Canonical Wnt Pathway, KEGG WNT Annotation
<i>CDH1</i>	NM_004360.2	-0.306	0.756	0.86	Adhesion, Development & Differentiation, EMTMetastasis
<i>EGFR</i>	NM_201282.1	0.838	0.757	0.86	Adhesion, Calcium Binding and Signaling, Cell Cycle, Development & Differentiation, Migration
<i>TGFB1</i>	NM_000660.3	0.0283	0.812	0.903	EMTMetastasis
<i>WNT5B</i>	NM_032642.2	-0.0997	0.813	0.903	Calcium Binding and Signaling, KEGG WNT Annotation
<i>FRZB</i>	NM_001463.2	0.174	0.818	0.903	WNT Signaling Negative Regulation
<i>CCND2</i>	NM_001759.2	-0.0898	0.84	0.915	Cell Cycle, Development & Differentiation, KEGG WNT Annotation, WNT Signaling Target Genes
<i>PPAP2B</i>	NM_003713.3	-0.259	0.856	0.926	Migration
<i>EGR1</i>	NM_001964.2	-0.149	0.881	0.928	Development & Differentiation, Transcription Factors

<i>TCF7</i>	NM_003202.2	0.0397	0.883	0.928	Canonical Wnt Pathway, KEGG WNT Annotation, Transcription Factors
<i>AHR</i>	NM_001621.3	-0.13	0.885	0.928	Cell Cycle, Transcription Factors
<i>VANGL1</i>	NM_138959.2	0.091	0.886	0.928	KEGG WNT Annotation
<i>BTRC</i>	NM_033637.2	-0.0186	0.887	0.928	KEGG WNT Annotation, WNT Signaling Negative Regulation, WNT Signaling Target Genes
<i>LRP6</i>	NM_002336.1	0.0948	0.904	0.939	Canonical Wnt Pathway, KEGG WNT Annotation, WNT Signaling Negative Regulation
<i>PRKX</i>	NM_005044.1	-0.0233	0.916	0.946	KEGG WNT Annotation
<i>RAC2</i>	NM_002872.3	-0.0121	0.928	0.951	KEGG WNT Annotation
<i>SFRP4</i>	NM_003014.2	-0.106	0.935	0.953	Canonical Wnt Pathway, KEGG WNT Annotation, WNT Signaling Negative Regulation
<i>STAT3</i>	NM_139276.2	-0.0278	0.943	0.955	EMTMetastasis
<i>CXCR4</i>	NM_003467.2	-0.0396	0.96	0.966	EMTMetastasis
<i>WNT3</i>	NM_030753.3	0.0117	0.985	0.985	Calcium Binding and Signaling, Canonical Wnt Pathway, KEGG WNT Annotation
<b>GENES BELOW THE BACKGROUND LEVEL</b>					
<i>CTGF</i>	NM_001901.2	<20 counts			
<i>PRKACG</i>	NM_002732.2	<20 counts			
<i>DPP10</i>	NM_020868.3	<20 counts			
<i>WNT1</i>	NM_005430.2	<20 counts			
<i>TWIST1</i>	NM_000474.3	<20 counts			
<i>FGF4</i>	NM_002007.2	<20 counts			
<i>MMP3</i>	NM_002422.3	<20 counts			

<i>IGF2</i>	NM_001127598.1	<20 counts			
<i>WNT8A</i>	NM_058244.2	<20 counts			
<i>PPP3R2</i>	NM_147180.2	<20 counts			
<i>TWIST2</i>	NM_057179.2	<20 counts			
<i>PROM1</i>	NM_006017.1	<20 counts			
<i>PLCB4</i>	NM_000933.3	<20 counts			
<i>WNT9A</i>	NM_003395.1	<20 counts			
<i>DKK2</i>	NM_014421.2	<20 counts			
<i>DKK1</i>	NM_012242.2	<20 counts			
<i>WISP1</i>	NM_080838.1	<20 counts			
<i>WNT8B</i>	NM_003393.2	<20 counts			
<i>FZD10</i>	NM_007197.2	<20 counts			
<i>FZD4</i>	NM_012193.2	<20 counts			
<i>WNT7A</i>	NM_004625.3	<20 counts			
<i>SOST</i>	NM_025237.2	<20 counts			
<i>KCNQ1</i>	NM_181798.1	<20 counts			
<i>SFRP2</i>	NM_003013.2	<20 counts			
<i>DLK1</i>	NM_003836.4	<20 counts			
<i>WNT16</i>	NM_057168.1	<20 counts			
<i>FGF7</i>	NM_004675.2	<20 counts			
<i>TLE1</i>	NM_005077.3	<20 counts			
<i>WIF1</i>	NM_007191.2	<20 counts			

<i>COL1A2</i>	NM_000089.3	<20 counts			
<i>DKK3</i>	NM_001018057.1	<20 counts			
<i>WNT9B</i>	NM_003396.1	<20 counts			
<i>FZD9</i>	NM_003508.2	<20 counts			
<i>PYGO1</i>	NM_015617.1	<20 counts			
<b>HOUSEKEEPING IN THE PANEL</b>					
<i>CC2D1B</i>	NM_032449.2	Housekeeping			
<i>COG7</i>	NM_153603.3	Housekeeping			
<i>EDC3</i>	NM_001142443.1	Housekeeping			
<i>GPATCH3</i>	NM_022078.2	Housekeeping			
<i>HDAC3</i>	NM_003883.2	Housekeeping			
<i>MTMR14</i>	NM_022485.3	Housekeeping			
<i>NUBP1</i>	NM_001278506.1	Housekeeping			
<i>PRPF38A</i>	NM_032864.3	Housekeeping			
<i>SAP130</i>	NM_024545.3	Housekeeping			
<i>SF3A3</i>	NM_006802.2	Housekeeping			
<i>TLK2</i>	NM_006852.2	Housekeeping			
<i>ZC3H14</i>	NM_001160103.1	Housekeeping			