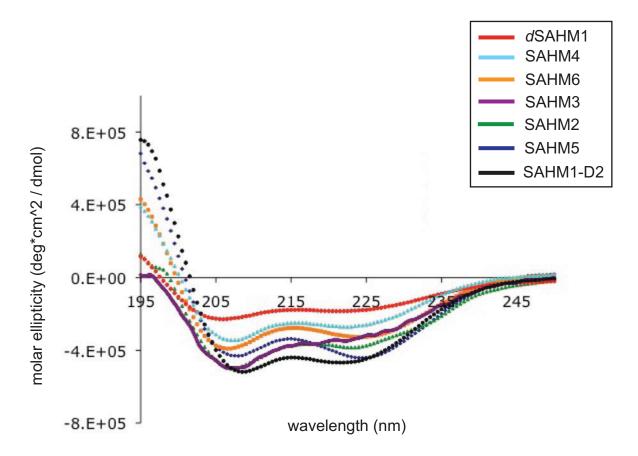
doi: 10.1038/nature08543

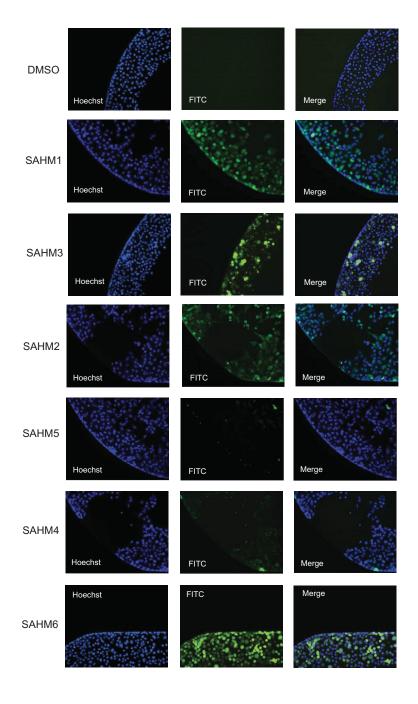
SUPPLEMENTARY INFORMATION

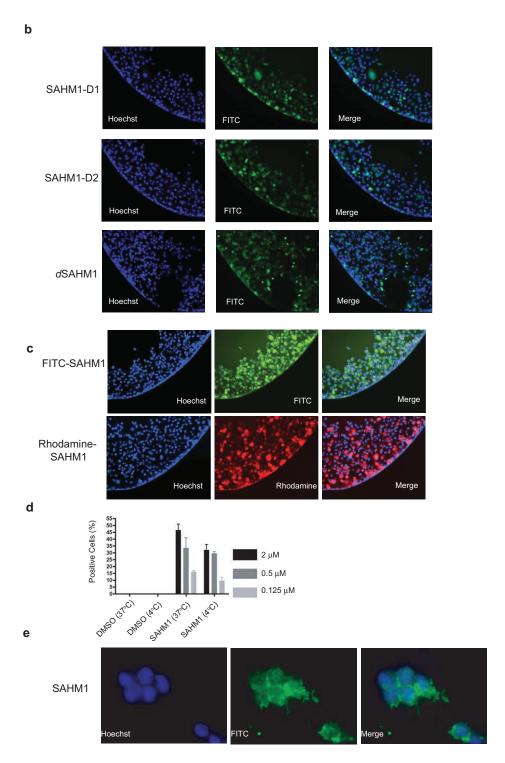


Supplementary Figure 1. CD spectra of SAHM peptides. Circular dichroism spectroscopy (CD) of additional MAML1 derived peptides and SAHM1 mutant peptides. Molar ellipticity at 222 nm was used to determine percent helicity at room temperature.

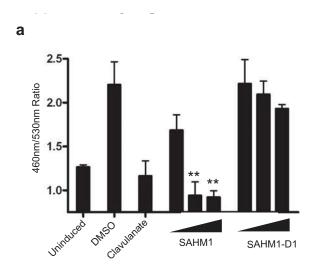
Supplementary Figure 2

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Supplementary Figure 2. SAHMs penetrate cells and access cytosolic and nuclear compartments. a) Analysis of FITC-SAHM peptide (5 μ M) intracellular access in U2OS cells by epifluorescent microscopy. After incubation for 12 hr, washed cells were imaged on Hoescht and FITC channels at 20X. b) Epifluorescence microscopy of the mutated peptides SAHM1-D1, SAHM1-D2 and *d*SAHM1, imaged as in (a). c) Intracellular penetration by rhodamine-labelled SAHM1, and FITC-SAHM1 imaged after 24 hr incubation. d) Temperature dependent uptake of FITC-SAHM1 after four-hour incubation at 37°C or 4°C. e) Confocal microscopy of FITC-SAHM1 treated KOPT-K1 cells (12 hr).





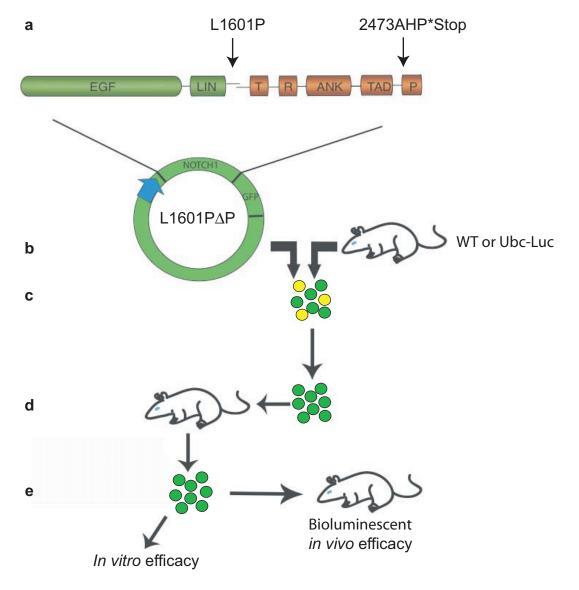
Supplementary Figure 3. Effects of SAHM peptides on NOTCH

transactivation. a) Suppression of the transactivation activity of ICN1 in HeLa cells by SAHM1, as assessed using a CSL-dependent β -lactamase reporter gene. Both SAHM1 and SAHM1-D1 peptides were dosed at 0.625, 2.5 and 10 μM, from left to right. Clavulanate (0.625μM) was used as a positive β -lactamase inhibitor control. **b)** Effect of vehicle (DMSO), DAPT (1μM) or SAHM1 (10 μM) on ICN1 protein levels in KOPT-K1 cells. Cells were treated for 24 hr prior to preparation of lysates, which were probed on Western blots with antibodies specific for ICN1 and GAPDH. All graphical representations represent mean \pm s.d. (n=3).

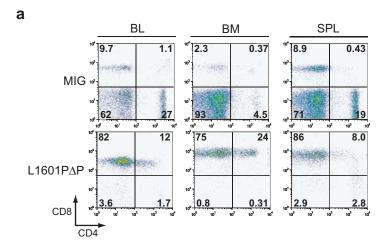
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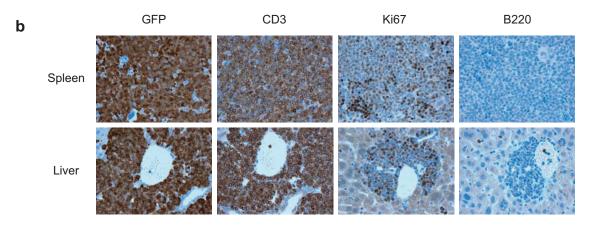
BROBE	CENE SYMBOL	DDODE	CENE CYMPOL
PROBE	GENE SYMBOL	PROBE	GENE SYMBOL
216557_x_at	IGHA1 /// IGHG1 ///	207515_s_at	POLR1C
228071_at	GIMAP7	219777_at	GIMAP6
219361_s_at	ISG20L1	214427_at	NOL1
218270_at	MRPL24	202613_at	CTPS
218597_s_at	ZCD1	203701_s_at	TRMT1
221712_s_at	WDR74	223018_at	NOB1
1564906_at	MATR3	222288_at	PPP4R2
218512_at	WDR12	203023_at	HSPC111
218866_s_at	POLR3K	201675_at	AKAP1
223413_s_at	LYAR	204133_at	RNU3IP2
201802_at	SLC29A1	222632_s_at	LZTFL1
224714_at	MKI67IP	218199_s_at	NOL6
203219_s_at	APRT	212456_at	KIAA0664
219006_at	C6ORF66	222875_at	DHX33
208799_at	PSMB5	227093_at	USP36
218105_s_at	MRPL4	218016_s_at	POLR3E
220865_s_at	PDSS1	218670_at	PUS1
209760_at	KIAA0922	203238_s_at	NOTCH3
204514_at	DPH2	203119_at	CCDC86
226410_at	LOC348180	218594_at	HEATR1
211949_s_at	NOLC1	210463_x_at	TRMT1
215792_s_at	DNAJC11	217884_at	NAT10
218238_at	GTPBP4	224632_at	GPATC4
217838_s_at	EVL	221649_s_at	PPAN
200895_s_at	FKBP4	217850_at	GNL3
219083_at	SHQ1	220688_s_at	C1ORF33
202138_x_at	JTV1	203612_at	BYSL
201489_at	PPIF	209100_at	IFRD2
225866_at	BXDC1	210448_s_at	P2RX5
201574_at	ETF1	211576_s_at	SLC19A1
232154_at	LOC199800	224634_at	GPATC4
216397_s_at	BOP1 /// LOC653119	209336_at	PWP2H
224596_at	SLC44A1	202431_s_at	MYC
214011_s_at	HSPC111	203737_s_at	PPRC1
209971_x_at	JTV1	218984_at	PUS7
201478_s_at	DKC1	225943_at	NLN
218758_s_at	D21S2056E	203394_s_at	HES1
63009_at	SHQ1	202934_at	HK2
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225682_s_at	POLR3H	227347_x_at	HES4

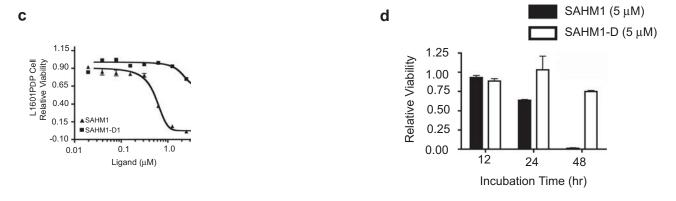
Supplementary Figure 4: GSEA analysis of SAHM gene expression profile and pathway specificity. a) Probe identifiers and gene names for the GSI-NOTCH gene set used in GSEA.



Supplementary Figure 5: Generation of the L1601PΔP & Luc: L1601PΔP models of NOTCH1 activation-dependent T-ALL. a) Schematic of the retroviral construct expressing a constitutively active *NOTCH1* allele bearing dual mutations initially identified in human T-ALL. EGF, EGF-like repeats; LIN, LIN-12 repeats; T, transmembrane domain; R, RAM domain; ANK, ankyrin repeat domain; TAD, transactivation domain; P, PEST regulatory domain. The relative locations of extracellular and PEST domain mutations are indicated. b) Bone marrow harvested from 5-fluorouracil treated adult wild-type or Ubc-Luc C57BL/6-Tyr^{C/C} mice was transduced with L1601PΔP vector. c) Isolated GFP+ cells were transplanted into primary recipient mice and followed for signs of morbidity, organomegaly and progressively increasing bioluminescence. d-e) Primary cells were collected from leukemic recipient animals and used for *in vitro* and *in vivo* studies.



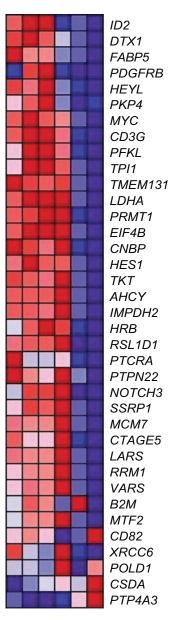




Supplementary Figure 6. Analysis of primary leukemic animals from the

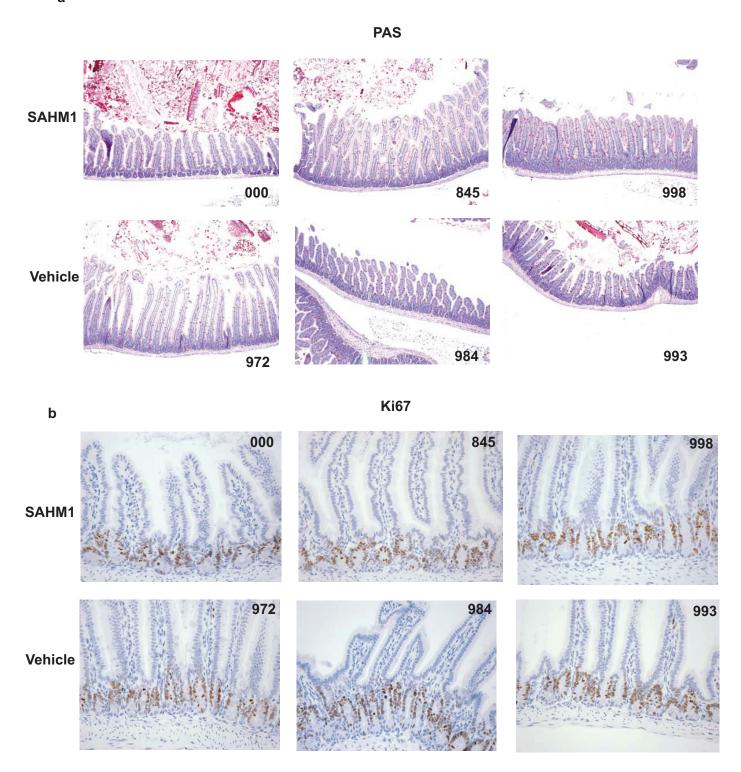
L1601PΔP T-ALL model. a) Flow cytometric analysis of tissues derived from primary leukemic L1601PΔP (donor) mice. b) Immunohistochemical staining of spleen and liver from primary leukemic (donor) mice. Both tissues are heavily infiltrated by GFP+/CD3+ blasts that are negative for the B cell marker B220. A high fraction of cells are also positive for Ki67, a cell cycle marker. c) Effects of SAHMs on primary murine T-ALL cells *in vitro*. Cell viability was measured using Cell Titer-Glo. d) Kinetic study of SAHM1 or SAHM1-D1 (5 μ M) treatment in sorted (FACS) primary leukaemia cells (GFP+CD3+) for 12, 24 or 48 hours. No impairment in viability is observed at 12 hours, establishing the duration of *ex vivo* SAHM1 exposure in studies of leukaemia initiation in secondary recipients.

Vehicle SAHM1



Supplementary Figure 7. GSEA analysis of *in vivo* gene expression profile in SAHM1 treated mice. a) The rank-ordered GSI/dnMAML1 gene-set curated from a published study in murine T6E cells, which is a T-ALL cell line dependent on mutant NOTCH1. This gene set was queried against the gene expression profile generated from SAHM1 (30mg/kg, bid, n=3) and vehicle treated mice (n=3) as discussed in Methods and presented in figure 5.

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Supplementary Figure 8. Effects of SAHM1 on gastrointestinal tract of treated mice. Gastrointestinal tracts from mice treated with SAHM1 (30mg/kg, bid, n = 3) or vehicle alone (n = 3) for five days were harvested and stained for a) PAS and b) Ki67 to assess proliferative changes and goblet cells. No apparent

changes were present.

Cell Line	NOTCH1 HD	NOTCH1 PEST	FBXW7	PTEN
КОРТ-К1	L1601P	2515 RVP*STOP	WT	WT
HPB-ALL	L1575P	Insertion 2442: EGRGRCSHWAPAAWRCTL FCPRRAPPCPRRCHPRWSP*STOP	WT	WT
DND-41	L1594P	Insertion 2444: CCSHWAPAAWRCTLFCPRR APPCPRRCHPRWSHP*STOP	WT	WT
TALL-1	WT	WT	WT	WT
CUTTL1	t(7;9): TCRB- NOTCH1 fusion transcript	WT	WT	WT
SUTP1	t(7;9): TCRB- NOTCH1 fusion transcript	WT	WT	WT
MOLT-4	L1601P	2515 RVP*STOP	WT	MUT
JURKAT	WT	WT	R505C	MUT
K-562 (BCR-Abl driven)	-	-	-	-

Supplementary Table 1. Genetic annotation of human T-ALL cell lines.

Mutational status of *NOTCH1*, *FBXW7* and *PTEN* in human T-ALL cell lines, curated from published reports. Notch mutational status imperfectly correlates with sensitivity to GSI. For example, T-ALL1 is a human cell line with wild-type *NOTCH1* widely reported as GSI-sensitive^{14,15,41,43}. At present, the mechanism of GSI-sensitivity is not known.

Supplemental Table 2

SSHNOTCH	NAME	SIZE	ES	NES	NOM p-val	FDR q-val	FWER p-val
VSHIFT (Q3							0
VSMAX_01							
VSNMYC 01 224 -0.423833 -1.8250479 0 0.009299679 0.042 GGAANGGGAANY UNKNOWN 92 -0.473845 -1.81350770 0 0.009429279 0.045 VSPAX5 02 15 -0.880415 -1.7850482 0.0056777 0 0.03889474 0.255 VSAP1 04 239 -0.383013 -1.6770188 0 0.03889474 0.225 VSAP1 04 239 -0.383013 -1.6770188 0 0.03456724 0.275 VSAP1 04 239 -0.383013 -1.6770188 0 0.03456724 0.275 VSEMYCMAX 02 226 -0.384965 -1.6554989 0 0.04255436 0 0.325 VSECFDDE1 01 141 -0.404592 -1.6551933 0 0.03871824 0 0.227 VSEZF DD1 01 195 -0.384104 -1.6467547 0 0.03719344 0 0.325 VSEZF DD1 01 195 -0.384104 -1.6467547 0 0.03714745 0 3.35 VSEAP 05 5 184 -0.385666 -1.640435 0 0.03312251 0 <t< td=""><td>_</td><td></td><td></td><td></td><td>_</td><td></td><td></td></t<>	_				_		
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VSAPT Q4	· <u> </u>						
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SGCGSSAAA_VSEZF10P2_01							
MCAATINNINNINGCG UIKKNÖWN							
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V\$USF_Q6_01							
V\$SRF_01 48 -0.413747 -1.440305 0.03358779 0.09008743 0.991							
V\$AP1 Q2 237 -0.327919 -1.4368657 0.00398936 0.09255326 0.993	V\$AP1_Q2		-0.327919			0.09255326	0.993

V\$ZF5 01	I 196	-0.336744	-1 4358346	0.01209677	0.09213992	0.994
V\$EGR Q6		-0.328971		0.00782269	0.091004595	0.994
V\$SP1 Q4 01	208			0.00702203	0.091779076	0.994
V\$ELF1 Q6		-0.340333		0.00900901	0.093245514	0.994
V\$ETS1 B		-0.332021		0.00923920	0.093243314	0.996
V\$AP1 Q6		-0.328669		0.01041667	0.09279920	0.996
V\$PEA3 Q6	198			0.01041007	0.09316525	0.998
V\$AP1FJ_Q2	240			0.00777202	0.092298985	0.998
WWTAAGGC UNKNOWN		-0.355822		0.00777202	0.092296965	
V\$SP1_Q6_01						0.998
		-0.334917		0.00662252	0.09028937	0.998
V\$E2F_Q6		-0.333851		0.01069519	0.09420662	0.998
V\$HNF4_01_B		-0.332195		0.00655308	0.0941269	0.998
CGGAARNGGCNG_UNKNOWN		-0.431889	-1.4149487		0.097995654	0.999
V\$E2F_Q4		-0.333696		0.0135318	0.09782387	0.999
V\$AP2_Q3		-0.328188		0.00780234	0.10858628	0.999
V\$PAX4_01		-0.323428		0.00762389	0.10738702	0.999
V\$E2F1DP1RB_01		-0.330201		0.01062417	0.10621905	0.999
V\$DR1_Q3		-0.327628		0.01058201	0.105627164	0.999
V\$TEL2_Q6	181	-0.335269	-1.3999805	0.01604278	0.1059651	0.999
V\$HNF4_DR1_Q3	214	-0.32134	-1.3987306	0.00903226	0.10578115	1
GCGNNANTTCC UNKNOWN	98	-0.356648	-1.3969787	0.03448276	0.106241964	1
V\$LFA1 Q6		-0.328002		0.01174935	0.108693734	1
V\$CMYB 01		-0.326078		0.00909091	0.10938369	1
CAGNYGKNAAA UNKNOWN	59			0.05247377	0.10990368	1
TMTCGCGANR UNKNOWN		-0.351593		0.02124646	0.109694675	1
TTCYRGAA UNKNOWN		-0.314163		0.00770218	0.11055498	1
TNCATNTCCYR UNKNOWN		-0.357059		0.03851852	0.11357373	1
GCTNWTTGK UNKNOWN		-0.337039				1
_ · · · · · · · · · · · · · · · · · · ·				0.00376412	0.11391955	
V\$PPAR_DR1_Q2		-0.319056		0.01456954	0.11903341	1
V\$E2F_Q6_01		-0.321567		0.02300406	0.119736485	1
V\$PPARG_01		-0.419125		0.07914764	0.120632745	1
V\$STAT1_01	54			0.07763975	0.124155946	1
V\$XBP1_01		-0.342906		0.03429355	0.124341704	1
V\$ATF4_Q2		-0.319577		0.01236264	0.123578146	1
V\$NRF2_Q4		-0.316891		0.01699347	0.12360512	1
V\$E2F1_Q4_01		-0.327054	-1.3675861	0.01583114	0.12268653	1
V\$STAT1_02	201	-0.321306	-1.3651437	0.0185923	0.12427513	1
V\$PAX4_03	226	-0.314403		0.01164295	0.12363862	1
V\$NRF1_Q6	198	-0.320746	-1.3642207	0.01886793	0.1228531	1
V\$NFE2_01	228	-0.314357	-1.3634562	0.0139417	0.122600205	1
V\$HNF4_01	219	-0.316886	-1.3626075	0.01237964	0.12235247	1
V\$STAT 01	215	-0.316323	-1.3605535	0.02533333	0.123995736	1
YGACNNYACAR UNKNOWN	79			0.05823864	0.12571198	1
V\$E2F Q3	180	-0.322517		0.03238867	0.12468991	1
V\$NERF Q2		-0.320299		0.01978892	0.124808684	1
V\$ATF 01		-0.320233		0.01643489	0.12458347	1
V\$SP1 01		-0.316821		0.01839685	0.12737526	1
TGASTMAGC V\$NFE2 01		-0.325235		0.01839083	0.12743075	1
V\$AREB6 04		-0.325235		0.03351206	0.12743075	1
V\$COUP_DR1_Q6		-0.318955		0.01864181	0.12604715	1
V\$ATF6_01		-0.346228		0.05988024	0.12704362	1
V\$SP1_Q2_01		-0.31705		0.02005348	0.12704669	1
V\$CHOP_01		-0.312523		0.02203857	0.14443153	1
V\$STAT3_01		-0.465608		0.13207547	0.14779325	1
V\$STAT1_03		-0.306527		0.02734375	0.1532541	1
AACWWCAANK_UNKNOWN		-0.336711		0.05807366	0.15293744	1
V\$CREB_Q2		-0.304496		0.02384106	0.15223567	1
KMCATNNWGGA_UNKNOWN	73	-0.354726	-1.3258634	0.07110439	0.15321198	1
V\$USF_02	225	-0.306839	-1.323693	0.03631647	0.15518682	1
V\$E2F Q3 01		-0.311845		0.0331565	0.1594396	1
WCAANNNYCAG UNKNOWN		-0.306316		0.03355705	0.1656129	1
V\$CREB 02		-0.305926		0.03178808	0.17347085	1
GGCKCATGS UNKNOWN		-0.375183		0.11520737	0.1751147	1
V\$PXR Q2		-0.301923		0.02486911	0.18112238	1
GGCNKCCATNK_UNKNOWN		-0.339108		0.07819549	0.18107131	1
OCCURROOMINITEDINININOVIN	<u>1</u> 94	0.000 100	- 1.0000100	0.07013043	0.1010/131	I.

NAVE US	200	U 3U34001	1 20220E2	0.04202477	0.10022409	4
V\$ARNT_02 V\$TGIF 01		-0.303188 -0.305351		0.04302477 0.04617414	0.19922498 0.20026325	1
V\$TATA 01		-0.299162		0.04617414	0.20026325	1
V\$HNF4 Q6		-0.299102		0.05345502	0.20076631	1
V\$SOX9 B1		-0.299881		0.04666667	0.20146428	1
V\$SRF C						
_		-0.303764		0.0646438	0.21600312	1
V\$MAZ_Q6		-0.307867	-1.2735022		0.22578734	1
V\$ETS_Q4		-0.296655		0.04842932	0.2271866	1
YTTCCNNNGGAMR_UNKNOWN		-0.399176		0.14772727	0.22684057	1
RGAGGAARY_V\$PU1_Q6		-0.275483		0.02696079	0.22839169	1
CGTSACG_V\$PAX3_B		-0.317643		0.08643815	0.2300422	1
V\$SREBP_Q3		-0.295237		0.05733333		1
GTTNYYNNGGTNA_UNKNOWN		-0.346653		0.11377245	0.23211029	1
RNCTGNYNRNCTGNY_UNKNOWN		-0.355264		0.11544462	0.23552467	1
V\$CREBP1CJUN_01		-0.291292		0.05269923	0.23646465	1
V\$SP3_Q3		-0.295189		0.06684857	0.24090566	1
V\$BACH2_01		-0.291876		0.06169666		
YRTCANNRCGC_UNKNOWN		-0.363755		0.14331724	0.24856639	1
V\$COUP_01	218	-0.288535		0.06468305	0.25128898	1
V\$AP2_Q6_01		-0.288754		0.08016305	0.25132385	1
V\$FXR_IR1_Q6		-0.318954		0.1038961	0.25145793	1
ACTWSNACTNY_UNKNOWN	73	-0.340481	-1.2472546	0.13394216	0.2561748	1
GTGGGTGK_UNKNOWN	245	-0.286134		0.05822785		1
V\$GCM_Q2	196	-0.292918	-1.2468034	0.06905711	0.25382936	1
V\$E2F_Q2	132	-0.307838	-1.2436256	0.08251748	0.25933644	1
V\$ICSBP_Q6	209	-0.289435	-1.2426231	0.072	0.25981545	1
V\$CREB_01	215	-0.28749	-1.240451	0.07105943	0.26336452	1
V\$IRF2_01	105	-0.316236	-1.2377642	0.12089552	0.26734734	1
V\$MZF1_02	195	-0.290323	-1.2371988	0.07662338	0.2670056	1
TGACGTCA V\$ATF3 Q6	190	-0.291008		0.06629834	0.2719174	1
V\$AP1 C	245	-0.28142	-1.2296448	0.06623377	0.28154048	1
YTCCCRNNAGGY UNKNOWN	60	-0.345175		0.14728682	0.2811707	1
V\$CETS1P54 01		-0.286992		0.07984293	0.28132096	1
V\$SMAD Q6	215	-0.284602	-1.2262139	0.08168643	0.28403303	1
V\$CREB_Q4		-0.287596		0.09043928	0.2835709	1
V\$LMO2COM 01		-0.285337		0.08481532	0.28196022	1
TGACCTTG_V\$SF1_Q6		-0.285034		0.08255659	0.2819899	1
V\$HSF Q6		-0.295066		0.09861111	0.282154	
V\$AP1 Q6 01		-0.283833		0.09736842	0.2852732	1
V\$E2F1_Q4		-0.286043		0.0923277	0.28753537	1
V\$PAX3 01		-0.458374		0.20973155	0.28589842	1
V\$SPZ1 01		-0.283189		0.10131579	0.29495767	1
V\$VDR Q6		-0.283362		0.09299363	0.2954499	1
CCCNNGGGAR_V\$OLF1_01		-0.274613		0.08035714	0.29459113	
V\$CREB_Q2_01		-0.289081		0.10225764		
V\$YY1 02		-0.286771		0.09917355	0.30269626	
V\$E4BP4_01		-0.284854		0.03317333	0.30139422	1
V\$SF1 Q6	206			0.10416666	0.3038719	
YRCCAKNNGNCGC UNKNOWN		-0.339793		0.1770335	0.3036718	
GGCNNMSMYNTTG UNKNOWN		-0.339793		0.1770333	0.30533102	
				0.17269719		
V\$NGFIC_01		-0.280537			0.30521166	1
V\$IRF_Q6		-0.283856		0.1171875	0.31707937	
V\$AP2_Q6		-0.27688		0.10303831	0.31550574	
V\$ER_Q6_01		-0.276968		0.11182519	0.3155869	
V\$SREBP1_Q6		-0.277074		0.10939511	0.3142548	
V\$AHRARNT_02		-0.44909		0.25127333	0.31633478	
GATGKMRGCG_UNKNOWN		-0.346597	-1.1949217		0.3214155	
TGTYNNNNRGCARM_UNKNOWN		-0.328046		0.17546584	0.3229151	1
V\$HEN1_02		-0.288455		0.13399154	0.3240815	
V\$E4F1_Q6		-0.275471		0.11914324	0.34306952	
RGAANNTTC_V\$HSF1_01		-0.261109		0.10278114	0.3438885	1
V\$SP1_Q6		-0.276417		0.12433155	0.36097425	
V\$MMEF2_Q6	234	-0.272075	-1.1762358		0.36134884	
YAATNRNNNYNATT_UNKNOWN	91	-0.30802	-1.1730983	0.18063584	0.3683257	1
GCGSCMNTTT UNKNOWN	56	-0.333953	-1.1728667	0.215478	0.3670974	1

V\$E2F1 Q3 01	208	-0.274217	-1.1671201	0.15314136	0.3816793	1
TGCTGAY UNKNOWN	455	-0.255391	-1.1664044	0.08232445	0.38173133	1
V\$MZF1_01		-0.271473	-1.1659796	0.14686248	0.38094485	1
V\$NFAT Q4 01		-0.269644		0.16688742	0.38479018	1
V\$CREB Q3		-0.271148		0.14557824	0.38607877	1
V\$CACCCBINDINGFACTOR Q6		-0.271345		0.16154873	0.38476664	1
V\$PR 01		-0.289081		0.18813314		1
V\$RFX1_01		-0.270012		0.16842106	0.4055444	1
V\$STAT3 02		-0.291955		0.19765739	0.4068542	1
V\$AREB6 03		-0.271098		0.17173052	0.41438282	1
V\$PAX3 B		-0.307133		0.22288261	0.41632634	
V\$HNF4ALPHA Q6		-0.265958		0.19505852	0.4360818	
V\$ER Q6		-0.266575		0.19303032		1
V\$NFY 01		-0.266912		0.10773033	0.43986043	1
V\$LEF1 Q2		-0.270599		0.19319728	0.44225582	1
V\$SREBP1 01		-0.276023		0.19662921	0.44223362	1
					0.45022014	
TGCGCANK_UNKNOWN		-0.247917		0.15658364		1
TGGNNNNNKCCAR_UNKNOWN	350			0.16730037	0.45213866	1
V\$HTF_01		-0.318882		0.25609756	0.4507304	1
RACCACAR_V\$AML_Q6		-0.262755		0.18915343	0.4522425	1
SYATTGTG_UNKNOWN		-0.264852		0.20476858		1
V\$AR_03		-0.331448		0.26564884	0.45985165	1
V\$MAZR_01		-0.268021		0.20603675	0.4627701	1
V\$AP1_Q4_01	225			0.21073298		1
V\$LEF1_Q6	-	-0.260625		0.22976501	0.4866212	1
V\$TFIIA_Q6		-0.259312		0.24057217	0.4849089	1
TTCYNRGAA_V\$STAT5B_01	272	-0.252876		0.21778351	0.4867588	1
V\$MYOGENIN_Q6		-0.259601		0.24898511	0.49721432	1
GTCNYYATGR_UNKNOWN	91	-0.289545	-1.1115562	0.27219796	0.49760178	1
CRGAARNNNNCGA_UNKNOWN	38	-0.335778	-1.1102667	0.28548896	0.49986416	1
V\$HEN1_01	165	-0.265629	-1.1082169	0.2465181	0.504769	1
TCANNTGAY_V\$SREBP1_01	392	-0.241108	-1.106247	0.19976498	0.50885814	1
V\$HMX1_01	38	-0.33436	-1.1009536	0.3176661	0.52451783	1
V\$NFKB_C	231	-0.255401	-1.098669	0.24577373	0.5300469	1
WCTCNATGGY_UNKNOWN	61	-0.305973	-1.0982162	0.3131783	0.52921146	1
V\$IRF7 01	212	-0.255168	-1.0971407	0.25695366	0.53057283	1
V\$PAX Q6		-0.254178		0.25751635		1
V\$TTF1 Q6	215	-0.256631		0.2621871	0.5321351	1
V\$CEBPB 02		-0.253048		0.27792206	0.5519305	1
V\$NFY Q6 01		-0.251152		0.28165373		1
V\$ETS2 B		-0.254097		0.28795812	0.5618426	1
RACTNNRTTTNC_UNKNOWN		-0.272789		0.29760227	0.5683926	1
V\$CDPCR3 01		-0.321513		0.35117057	0.56713015	1
V\$E47 02		-0.254642		0.2948886	0.576616	1
CCTNTMAGA UNKNOWN		-0.271058		0.3069307	0.57648885	1
V\$MYOD Q6		-0.271038		0.3009307		
V\$MTF1 Q4		-0.252503		0.29267596	0.5763471	1 1
CATTGTYY V\$SOX9 B1		-0.251665		0.2996159	0.5893229	1
V\$EGR1_01		-0.242259		0.2914573	0.59325916	1
V\$EGR1_01 V\$E2A Q2				0.28074867		1
V\$EZA_QZ V\$PAX8_B		-0.248358			0.5988252	
		-0.280808		0.32544377	0.59705466	1
V\$CEBPA_01		-0.250207		0.29419526	0.6019033	1
V\$TFIII_Q6		-0.252512		0.3263434	0.605123	1
V\$AP1_01		-0.244849		0.33117723	0.61991626	1
V\$HLF_01		-0.245514		0.33110815	0.62470484	1
V\$CP2_01		-0.246686		0.33022636	0.62462664	1
V\$AR_01		-0.264385		0.3598862	0.62829643	1
V\$T3R_Q6		-0.245675		0.33678755	0.62947476	1
V\$CREL_01		-0.244018		0.34214002	0.6273018	1
V\$CEBPB_01		-0.244709		0.32774192	0.62570095	1
V\$DR3_Q4		-0.264084		0.35561877	0.629439	1
V\$IRF1_01	207	-0.243319	-1.0498049	0.34954008	0.6362813	1
V\$MYOD 01				0.04040044	0.05400004	-
[ν φίνι 1 O D_ O 1	207	-0.244668	-1.044919	0.34912044	0.65130234	1
V\$ISRE 01		-0.244668 -0.248918		0.34912044	0.65130234	1

YYCATTCAWW UNKNOWN	160	-0.252349	-1 0439423	0.35309973	0.64716	1
ARGGGTTAA UNKNOWN		-0.266251		0.3810219	0.65774226	1
V\$NKX62 Q2		-0.245822		0.36437246	0.6574888	1
V\$OLF1 01		-0.239956		0.36903226	0.6589575	1
V\$AP2ALPHA 01		-0.242515		0.36340207	0.6613056	1
V\$NKX25 01		-0.262105		0.38864627	0.66015816	1
V\$MAF Q6		-0.241047		0.36617842	0.6610602	1
V\$SOX5 01		-0.239173		0.37209302	0.6600091	1
V\$AP2REP 01		-0.250545		0.35944057	0.6636111	1
V\$AML Q6		-0.230345		0.37967914	0.67184895	1
V\$SRF Q5 01		-0.242093		0.39005235	0.67068475	1
		-0.241949				
WTTGKCTG_UNKNOWN				0.38210398	0.6710245	1
TAANNYSGCG_UNKNOWN		-0.287063		0.41252005	0.6741717	1
RYAAAKNNNNNTTGW_UNKNOWN		-0.277102		0.42065868	0.6752837	1
V\$COREBINDINGFACTOR_Q6		-0.237475	-1.0246178		0.68202484	1
V\$DR4_Q2		-0.237539		0.40832248	0.685572	1
V\$SRF_Q6		-0.236643		0.41192052	0.6948517	1
V\$BACH1_01		-0.234085		0.4213264	0.6952216	1
GTTRYCATRR_UNKNOWN		-0.252653		0.4119318	0.69664943	1
V\$STAT5A_01	211	-0.23691		0.41153845	0.6996776	1
V\$ZIC2_01		-0.235876	-1.0161893		0.69792175	1
V\$STAT6_01	218			0.4365285	0.7075428	1
V\$CP2_02		-0.237743		0.43386245	0.7101147	1
V\$FOXO4_01		-0.236848		0.44113666	0.71237123	1
V\$SRY_02	212	-0.233546		0.4224806	0.71194243	1
V\$E12_Q6	199	-0.2372	-1.0085809	0.43678162	0.7126142	1
WGTTNNNNNAAA_UNKNOWN	466	-0.217984		0.43550295	0.7183216	1
V\$YY1_01	210	-0.234372	-1.0037863	0.44987145	0.724732	1
ATGGYGGA_UNKNOWN	79	-0.266818	-1.0002711	0.45870206	0.7349637	1
V\$P300_01	201	-0.234538	-0.99610263	0.44702843	0.74777126	1
V\$AR_Q2	101	-0.256148	-0.99194926	0.46814814	0.76009464	1
V\$ER_Q6_02	213	-0.230593	-0.99134064	0.48687664	0.7597649	1
V\$SMAD3 Q6	196	-0.23563	-0.9894759	0.47925034	0.76424813	1
V\$TEF1 Q6	178	-0.235167	-0.9853324	0.48724833	0.7771921	1
RYTAAWNNNTGAY UNKNOWN	58	-0.274785	-0.98372704	0.49846625	0.78026944	1
V\$CREBP1 01		-0.239596		0.49788433	0.7807154	1
V\$NFKB Q6		-0.226992	-0.98078835		0.7854151	1
V\$PPARA 02		-0.251617	-0.98013437		0.78517807	1
V\$SRY 01		-0.234578	-0.9798824		0.7835038	1
V\$SMAD4 Q6		-0.230873	-0.97895676		0.7841558	1
V\$CEBP C		-0.232262	-0.9763328		0.7907677	1
ACCTGTTG UNKNOWN		-0.245033			0.7956259	1
RRAGTTGT UNKNOWN		-0.226346		0.51660025	0.79839456	1
CCAWWNAAGG V\$SRF Q4		-0.261266			0.80728203	1
V\$CREBP1_Q2		-0.227344		0.53350854		
V\$OSF2 Q6		-0.224882			0.8108521	1
V\$TBP 01		-0.226211	-0.96707237		0.8114685	1
V\$MYOD Q6 01		-0.226211	-0.96556854		0.810974	
V\$PAX8_01		-0.312243		0.51480263	0.81736916	
V\$SRF_Q4		-0.224379			0.8259001	1
V\$VDR_Q3		-0.223601	-0.95940614		0.82458425	1
V\$GR_01		-0.228367		0.5506849	0.82329714	
RYCACNNRNNRNCAG_UNKNOWN		-0.26748		0.52194357	0.8240381	1
V\$AP4_01		-0.219993			0.8229928	
V\$LBP1_Q6		-0.228134		0.55186725	0.82079774	
V\$HAND1E47_01		-0.221877		0.57198954	0.8218703	1
TGAYRTCA_V\$ATF3_Q6		-0.208306	-0.95564884		0.8220794	1
V\$STAT5B_01		-0.223562		0.57563585	0.8204764	1
V\$CREB_Q4_01		-0.226778			0.8207502	1
V\$P53_02		-0.219903		0.59395534	0.83230627	1
V\$ATF_B		-0.228845	-0.94790155	0.5635135	0.83854705	1
TTOOMOANA VACCEDED OO		-0.26917	-0.9470659	0.5421133	0.8388174	1
TTGCWCAAY_V\$CEBPB_02			-0.947 0039	0.000	0.0000	
V\$NFY_Q6		-0.220003	-0.94537455		0.84229594	1
	212			0.6026667		1

GGAMTNNNNNTCCY UNKNOWN	96	-0.239763	-0.94299203	0.54232424	0.8426266	1
V\$AP2GAMMA 01		-0.219834			0.84170127	1
TTCNRGNNNNTTC V\$HSF Q6	125	-0.235121	-0.9412346	0.57954544	0.8434613	1
V\$HSF1_01	218	-0.219177	-0.94020647	0.6051613	0.8443664	1
V\$HEB_Q6	222	-0.217064	-0.9368511	0.6475943	0.8532238	1
V\$CEBP_Q3	205	-0.218008	-0.9355038	0.61738	0.8550053	1
V\$CACBINDINGPROTEIN_Q6	198	-0.220421	-0.9339569	0.63502675	0.8577542	1
V\$ERR1_Q2	221	-0.217469	-0.93273294	0.6231691	0.8593525	1
V\$TAL1ALPHAE47_01	211	-0.215777	-0.9271164	0.64257556	0.87540174	1
V\$PAX2_02	214	-0.216499	-0.92594653	0.6323155	0.87672013	1
V\$GFI1_01	223	-0.212974	-0.92592627	0.65929776	0.8741645	1
V\$ATF1_Q6	191		-0.9235589	0.6586667	0.8792025	1
V\$TAXCREB_01		-0.227989	-0.922692	0.6396011	0.87939197	1
V\$AML1_01	207	-0.215321	-0.92197055	0.64470285	0.8791895	1
V\$EGR2_01		-0.220371		0.6586667	0.88274074	1
V\$GR_Q6		-0.215366	-0.9183374	0.6864516	0.88584286	1
YGCANTGCR_UNKNOWN		-0.234532		0.60926193	0.8860671	1
V\$HSF2_01		-0.212543			0.88364136	1
V\$EFC_Q6		-0.212075		0.66753244	0.88738066	1
V\$FOXO4_02		-0.211853		0.6626984	0.8868118	1
V\$AML1_Q6		-0.215321	-0.91386694		0.8873734	1
V\$MIF1_01		-0.221585	-0.91220284		0.890085	1
V\$TCF11MAFG_01		-0.21923		0.66050196	0.8894926	1
V\$STAT4_01		-0.210311		0.6939314	0.8905731	1
V\$CEBP_Q2		-0.214084	-0.90930146		0.89147466	1
V\$HNF3ALPHA_Q6		-0.214834	-0.9072031		0.89529777	1
CCAATNNSNNNGCG_UNKNOWN		-0.260893			0.8953056	1
V\$FAC1_01		-0.216061		0.6975477	0.89739615	1
TAAYNRNNTCC_UNKNOWN		-0.21927		0.6924138	0.9066092	1
V\$FOX_Q2		-0.212608			0.9062085	1
YNGTTNNNATT_UNKNOWN	292		-0.89942414		0.90694183	1
V\$FREAC3_01 V\$PTF1BETA_Q6		-0.210714	-0.89928406		0.9047491 0.91152006	1
CCAWNWWNNNGGC UNKNOWN		-0.211319 -0.242798	-0.89624643 -0.8935208		0.9169907	1
V\$PBX1 02		-0.242796		0.70679885	0.937512	1
V\$HNF3B 01		-0.220316	-0.88282263		0.94325763	1
V\$IPF1 Q4		-0.207744	-0.87761354		0.9555579	1
CCCNNNNNAAGWT UNKNOWN		-0.231429			0.957084	1
TTANWNANTGGM UNKNOWN		-0.253063		0.6807818	0.95447004	1
V\$E47 01		-0.203156	-0.8680921		0.9744583	1
RYTGCNNRGNAAC V\$MIF1 01		-0.239045			0.97519016	1
V\$EVI1 06		-0.301442	-0.85762835		0.9977648	1
V\$RP58_01		-0.20338		0.79591835	0.99649286	1
V\$NFY C		-0.200848		0.8214748	0.99722594	1
V\$STAT Q6		-0.198919		0.8069948		1
V\$GATA1 01		-0.199808			1	1
CTAWWWATA_V\$RSRFC4_Q2		-0.191885			1	1
V\$NF1_Q6_01	229	-0.195311	-0.8467041	0.83825415	1	1
V\$STAT5A_03		-0.196547		0.87614083	1	1
V\$NF1_Q6	234	-0.195919		0.8470437	1	1
SNACANNNYSYAGA_UNKNOWN	63	-0.231503		0.72981364	1	1
MYAATNNNNNNNGGC_UNKNOWN		-0.214518		0.7877493	1	1
RYTGCNWTGGNR_UNKNOWN		-0.21812		0.76217765	1	1
V\$PBX1_01		-0.196729		0.86092716	1	1
V\$TITF1_Q3		-0.195038		0.8733509	1	1
TTAYRTAA_V\$E4BP4_01		-0.19412		0.88676673	1	1
V\$AP3_Q6		-0.192048	-0.83001465		1	1
V\$EGR3_01		-0.228159			1	1
V\$ALPHACP1_01		-0.193743			1	1
V\$CDPCR3HD_01		-0.194973	-0.82842517		1	1
V\$ATF3_Q6		-0.19297	-0.82820344		1	1
V\$GATA4_Q3		-0.193647	-0.82743406		1	1
AAGWWRNYGGC_UNKNOWN		-0.216509			1	1
V\$GATA1_05		-0.190597		0.87533873	1	1
V\$HFH1_01	205	-0.192582	-0.8205376	0.88962764	1	1

DTTTNINIVTCCM LINKNOWN	124	0.204276	0.9100525	0 0 4 6 0 0 0 5	1 1	1
RTTTNNNYTGGM_UNKNOWN V\$NFAT_Q6		-0.204276 -0.19295		0.8468085 0.875		<u>1</u> 1
V\$ARP1 01			-0.81750524			<u>1</u>
_ ·	231				1	
V\$IK2_01 V\$FREAC2_01					1	1
		-0.190102				1
V\$HFH3_01		-0.194889		0.87516963	1	1
CATRRAGC_UNKNOWN		-0.203602		0.86206895	1	1
V\$IRF1_Q6		-0.188359		0.9080311	1	1
V\$MEIS1_01		-0.189989			1	1
V\$HMGIY_Q6		-0.189197		0.9070385	1	1
GTTGNYNNRGNAAC_UNKNOWN		-0.212934			1	1
V\$FOXO1_02		-0.187048			1	1
V\$STAT5A_02		-0.20121			1	1
V\$CIZ_01			-0.79778075		1	1
V\$LYF1_01		-0.185133		0.9276486	1	1
V\$POU6F1_01		-0.186233		0.9351852	1	1
V\$RFX1_02		-0.185581		0.9247312	1	1
V\$RREB1_01	175	-0.189708			1	1
V\$TCF4_Q5		-0.185361			1	1
V\$TAXCREB_02	22	-0.270045		0.7641997	1	1
CCANNAGRKGGC_UNKNOWN	92				1	1
V\$RORA1_01	212	-0.181767		0.9401042	1	1
GGCNRNWCTTYS_UNKNOWN	64		-0.7799267	0.83753943	1	1
V\$SREBP1_02		-0.208007	-0.77747375	0.86158633	1	1
V\$IK1_01	224	-0.179157	-0.77448916	0.95454544	1	1
WGGAATGY_V\$TEF1_Q6	307	-0.173595	-0.7742006	0.9576923	1	1
V\$FOXJ2 02	193	-0.181948		0.94926566	1	1
V\$CDPCR1 01		-0.196415	-0.7697858	0.92395985	1	1
AGCYRWTTC_UNKNOWN		-0.197475		0.8949704	1	1
V\$PAX5 01		-0.190582			1	1
V\$TCF11 01	197	-0.17906		0.9546061	1	1
SMTTTTGT UNKNOWN		-0.169221	-0.76139516		1	1
V\$DBP Q6		-0.177197	-0.76000816		1	1
V\$AREB6 01		-0.176837	-0.75606203		1	1
V\$NFKAPPAB 01		-0.176567	-0.75475913		1	 1
V\$TATA C		-0.171903		0.9782609	1	 1
V\$NKX25 02		-0.174753			1	 1
V\$AR Q6		-0.172707		0.98271275	1	1
YKACATTT UNKNOWN		-0.170954		0.9896104	1	1
V\$WHN B		-0.173806		0.98300654	1	 1
V\$CDC5 01		-0.171665			1	<u>.</u> 1
YATTNATC UNKNOWN		-0.163767			1	 1
GGATTA_V\$PITX2_Q2		-0.156763	-0.7213298		1	1
V\$NFKAPPAB65 01		-0.167936			1	1
V\$LXR Q3		-0.201095		0.92762184		1
V\$AREB6 02		-0.201093				1
AAAYWAACM V\$HFH4 01		-0.164219				1
V\$HFH4 01		-0.166016				<u>1</u>
V\$AP1_Q2_01		-0.160114	-0.70512444			<u></u>
YCATTAA UNKNOWN		-0.15056			1	1
		-0.166152				<u> </u>
V\$PAX4_04		-0.166152		0.9896373		
V\$PPARA_01 V\$RSRFC4 Q2				0.90802675		<u> </u>
		-0.161135				
V\$GNCF_01		-0.191507		0.95510834		1
V\$GATA2_01		-0.179924		0.9776786		1
V\$GATA3_01		-0.159998				1
V\$AP4_Q6_01		-0.157946		0.99866843		1
V\$FOXD3_01		-0.159078				1
RNTCANNRNNYNATTW_UNKNOWN		-0.188924				1
V\$RSRFC4_01		-0.155671		0.9986807	1	1
V\$TAL1BETAE47_01		-0.154335		0.99866664		1
V\$FOXJ2_01		-0.158919		0.99728996		1
V\$CEBPDELTA_Q6		-0.155148				1
V\$AP4_Q5		-0.152364			1	1
V\$HNF3_Q6	164	-0.155125	-0.65333986	0.9958506	1	1

V\$ZIC3_01	217	-0.147387	-0.6324055	1	1	1
V\$RORA2_01	132	-0.155911	-0.6316755	0.99857146	1	1
V\$FREAC4_01	122	-0.15585	-0.625391	0.99559474	1	1
CTGRYYYNATT_UNKNOWN	55	-0.175999	-0.60784626	0.9823718	0.9996495	1
ACAWYAAAG_UNKNOWN	88	-0.150415	-0.5805317	0.99854016	0.9985612	1

Supplementary Table 2. Global transcription factor gene set analysis.

Results from GSEA analysis of all available transcription factor-target (TFT) gene sets in the Molecular Signatures Database applied to the SAHM1 expression profile in KOPT-K1 and HPB-ALL cells. From left to right, columns contain the gene set name from the MSigDB, the number of genes in the gene set, the overall enrichment score (ES), the normalized enrichment score (NES), the nominal p-value (NOM p-val), the false discovery rate q-value (FDR q-val), and the family-wise error rate p-value (FWER p-val). In general, gene sets with high positive or negative NES, low FDR q-value and low FWER p-value are considered to be significantly enriched. Gene sets have been ranked by the most stringent filter of FWER p-value. By any statistical measure, however, the GSI-NOTCH gene set is the most highly correlated with the SAHM1 expression profile.