

Supplementary figure legends

Figure S1: Preliminary immunophenotypic analysis prior to Cytobank export and patient cohort details

- A. Table illustrating the sample volume and percentage purity obtained after the first passage (Run1) and second passage (Run2) through the magnet during magnetic bead enrichment of CD34+ HSPCs. All samples had a purity of >95% prior to culture.
- B. Gating strategy on D0 prior to Cytobank import. Identification of singlets on forward scatter-area (FSC-A) versus (vs) FSC-height (H) plots was followed by the identification of cells on FSC-A vs side scatter-area (SSC-A), viable cells on SSC-A versus Viability and then the population of interest (POI) on FSC-A vs SSC-A. Zombie Violet (ZV) was used as the viability dye. Cells are coloured using banded colouring and plots are in a linear scale for all plots including C and D below.
- C. Gating strategy on D7 following the same method as mentioned in B above.
- D. Table displaying the characteristics of the HEU cohort. All mothers had a viral load (VL) <100 copies/mL within 3 months of delivery (D) (LTDL-Lower than detectable limit) and all except one were taking TLD as antiretrovirals (ARVs). One mother was on TEE (TDF, Emtricitabine and Efavirenz); Polymerase chain reaction (PCR) results for all infants were negative as was the HIV Gene Xpert on the umbilical cord blood (UCB).

Figure S2: Concatenated viSNE plots showing differences between HEU and HUU CD34+Lin- subsets

- A. Concatenated viSNE plot of D0 HUU samples coloured according to marker expression on the z-axis and scale of expression on the right of each channel. Only the expression of the Lin-CD34+ population is shown for the following markers: CD38 ECD, CD133 PE-Vio770, CD45RA APC, CD117 PE, CD90 BV510, and CD49f SB780 (N=4).
- B. Concatenated viSNE plot of D0 HEU samples with plot characteristics as described in A above (N=5).
- C. D7 viSNE concatenated samples for the HUU VC condition portraying the difference in immunophenotypic populations compared to the D0 samples. Cells cluster completely differently than for D0, and HSPC populations are predominantly CD38- on D7, compared to CD38+ on D0.
- D. Equivalent viSNE concatenated samples for the HEU VC portraying the differences compared to HUU. The HSPCs cluster differently. A large bright CD90+ population present in HUU samples is absent in HEU samples; however, HEU groups have

widespread dim expression of CD90 in the majority of cells. In contrast, HUU groups appear to have more dim expression of CD49f than HEU.

- E. FlowSOM clustering maps coloured according to the metaclusters (MCs), illustrated on the left of the image, for HUU and HEU samples (D0, D7 VC, D7 SR1). Only one sample is shown for ease of visualisation. Overt differences are the prominence of MC3 and 13 in HUU and MC 9 and 12 in HEU samples.
- F. Concatenated image of viSNE plot of D0 HUU samples showing only two channels (CD38-ECD and CD133 PE-Vio-770). The manual gates drawn around the cell clusters are shown with the names which were assigned to each population. (N=5) See also Table 1 for the phenotypic description of each population. Gates were guided by the FlowSOM MC in Figure S2E.
- G. Complimentary viSNE plot of D7 VC also only showing the two channels mentioned above in F. The plot clearly shows the marked difference in clustering between D0 and D7 samples. (N=5)

Fig. S3 Statistically significant metacluster differences in Lin-CD34+ sub-population between HEU and HUU populations.

- A. Box plot of the 17 FlowSOM MCs (x-axis) and frequency of each MC in HUU (orange) and HEU (blue) on D0. MC13, 15, and 17 were statistically different, with MC13 being higher in HUU and MC15 and 17 being higher in HEU. N=4 (HEU) and N=5 (HUU) for all experiments. *($p < 0,05$)
- B. Heatmap of the statistically significant MCs identified in A, portraying the phenotype of each MC. Scale of the plot is provided at the bottom of the image. MC13, is in keeping with ExtCMP.
- C. Box plot of the D7 VC for HEU and HUU populations illustrates the statistically significant MC3, 11 and 15. *($p < 0,05$)
- D. Corresponding heatmaps for C showing the phenotype of the MCs. MC3 corresponds to ExtLMPP2.
- E. Box plot of D7 SR1 for HEU and HUU populations. The MCs of significance are 2, 3, 4, 6, 11, and 12. *($p < 0,05$)
- F. The complementary heatmap for E shows the following correlation with phenotype. MC3 and 4 correspond to ExtLMPP2, MC2 to HSPC1 and 2 and MC12 to ExtLMPP1 and 3.

Figure S4: CITRUS cluster analysis showing differences between D0 and D7 HUU and HEU CD34+Lin- subsets

- A. Citrus analysis for D0 HUU vs HEU samples. Each CITRUS algorithm was repeated five times, using Significance Analysis of Microarrays (SAM) and abundance for each condition to obtain the common significant clusters. Equal events were used for each sample. Figure shows two of the five runs. Each circle depicts one cluster, and the red clusters are those found to be significantly different. The red clusters which give rise to other clusters are seen as the dominant clusters and those are used for export and phenotypic description. Significant MCs are 17935 (Lin-CD34+CD38+CD133dimCD45RA-CD117+CD90-CD49f-: correlates to ExtEMP2), 17936 (Lin-CD34+CD38+CD133+CD45RA-CD117+CD90-CD49f-: correlates to ExtEMP), 17953 (Lin-CD34+CD38dimCD133+CD45RA-CD117+CD90-CD49fdim-: correlates to ExtMPP), 17959 (Lin-CD34+CD38-CD133vCD45RA+CD117-CD90-CD49fv: correlates to ExtLMPP), and 17962 (Lin-CD34+CD38dimCD133+CD45RAvCD117+CD90-CD49f-: correlates to ExtCMP)
- B. Citrus analysis showing two sample runs for HUU D7 VC vs HEU D7 VC. Clusters 62995 (Lin-CD34+CD38-CD133+CD45RA+CD117vCD90-CD49f+: correlates to ExtLMPP1), and 62997 (Lin-CD34+CD38-CD133+CD45RA+CD117vCD90-CD49fv: correlates to ExtLMPP3)
- C. Similar analysis for HUU D7 SR1 vs HEU D7 SR1 samples is shown. Significant clusters are 269995 (Lin-CD34+CD38CD133v CD4RA+CD117dim/-CD90-CD49f+: correlates to ExtLMPP3) and 269996 (Lin-CD34+CD38-CD133+CD45RA+CD117vCD90dimCD49f-: correlates to ExtLMPP2).

Fig. S5 D0 CITRUS cluster immunophenotype analysis of Lin-CD34+ sub-population between HEU and HUU subgroups.

- A. Histogram plots of significant clusters identified during CITRUS analysis of D0, HUU (N=7) vs HEU (N=5). The blue shaded area depicts all the cells that were included in the algorithm and the red shaded area shows the histogram image of the specific cluster, and thus the phenotype. The immunophenotypic channel is shown above the figure and the cluster identification is on the right of each histogram.
- B. Box plot of the identified clusters of significance which illustrated the group within which the higher/lower event count (y-axis) was found (either HEU or HUU). On D0, 2 clusters (17953 and 17959, corresponding to ExtMPP and ExtLMPP respectively) were

more prominent in HEU samples and 3 in HUU (17935, 17936, 17962, ExtEMP2, ExtEMP and ExtCMP, respectively).

Figure S6: D7 CITRUS cluster immunophenotype analysis of Lin-CD34+ sub-population between HEU and HUU subgroups.

- A. Histograms of the two significant clusters identified in the D7 VC, HEU (N=4) vs HUU (N=5) samples from the CITRUS analysis. The CITRUS analysis was performed using the Significance Analysis of Microarrays (SAM) model in Abundance mode, with equal events across all subgroups. A false discovery rate (FDR) of 5% was applied to identify significant clusters, which were further visualised using cluster and box plot analyses. The blue shaded area depicts all cells included in the algorithm, while the red shaded area shows the histogram of the specific cluster, and thus the phenotype. The immunophenotypic channel is shown above the figure and the cluster identification is on the right side of each histogram.
- B. Box plot of exported significant CITRUS clusters on D7 VC, depicting clusters 62995 and 62997, corresponding to ExtLMPP1 and 3, found in a higher proportions in HEU samples.
- C. Histograms of the two significant clusters identified in the D7 SR1, HEU (N=4) vs HUU (N=5) CITRUS analysis.
- D. Box plot of significant CITRUS clusters, one which was higher in HEU samples (269995, corresponding to ExtLMPP3) and one in HUU samples (269996, corresponding to ExtLMPP2).

Figure S7: Concatenated UMAP plots showing differences between D0 and D7 HEU and HUU CD34-Lin- subsets

- A. UMAP image of HUU D0 Lin-CD34- population showing all the phenotypic markers above on the z-scale. Map is coloured by the scale of expression on the right. CD38-ECD, CD133 PE-Vio770, CD45RA APC, CD117 PE, CD90 BV510, CD49f SB 780 are shown (N=5). All events were included in the analysis.
- B. Complimentary UMAP portraying the HEU D0 population is shown with all the characteristics and channels as described above. (N=4). Visually the differences between HUU and HEU samples on D0 are not easily perceptible.

- C. Concatenated UMAP of HUU D7 VC with all features and channels as mentioned in A above. The D7 population analysis is not linked to the D0 analysis as the cells plated on D0 are a pure Lin-CD34+ population.
- D. UMAP of the HEU D7 VC population depicting easily perceptible changes as compared to HUU D7 VC. CD90 expression is much more widespread in HEU samples.
- E. The HUU D7 SR1 concatenated UMAP showing similar characteristics to the HUU D7 VC UMAP in B above.
- F. Complimentary HEU D7 SR1 UMAP confirming increased expression of CD90 in HEU samples.
- G. ViSNE plot showing the manual gates on an HUU samples guided by the FlowSOM MCs. Gate names are provided. Plot scale is on the right.
- H. FlowSOM MC dot overlays of D0, D7 VC, and D7 SR1 of one HUU sample showing the MCs which guided manual gates.

Figure S8: Population frequencies and statistically significant differences in Lin-CD34- sub-populations

- A. Table depicting the sub-population frequencies of the Lin-CD34- populations between HEU and HUU samples
- B. Bar graph of statistically significant MC in D7 VC. Orange bars are HEU and blue bars HUU samples. MC3, 13, and 16 are higher in HEU samples, while MC11 and 12 are higher in HUU samples.
- C. Corresponding bar graph of D7 SR1 conditions. MC13 and 16 are higher in HEU samples and MC6, 11, and 12 higher in HUU samples.

Figure S9: Differences in CFU phenotype between HEU and HUU samples.

- A. FlowSOM Dot overlay map showing the 17 identified MCs on the HUU samples, coloured according to the MC map. All 4 HUU samples are shown.
- B. Equivalent HEU FlowSOM map showing all 4 HEU samples. The MC confirm barely perceptible changes between sample groups
- C. FlowSOM MC box plot shows no significant difference in immunophenotype between the HEU and HUU sub-groups. *($p < 0.05$)

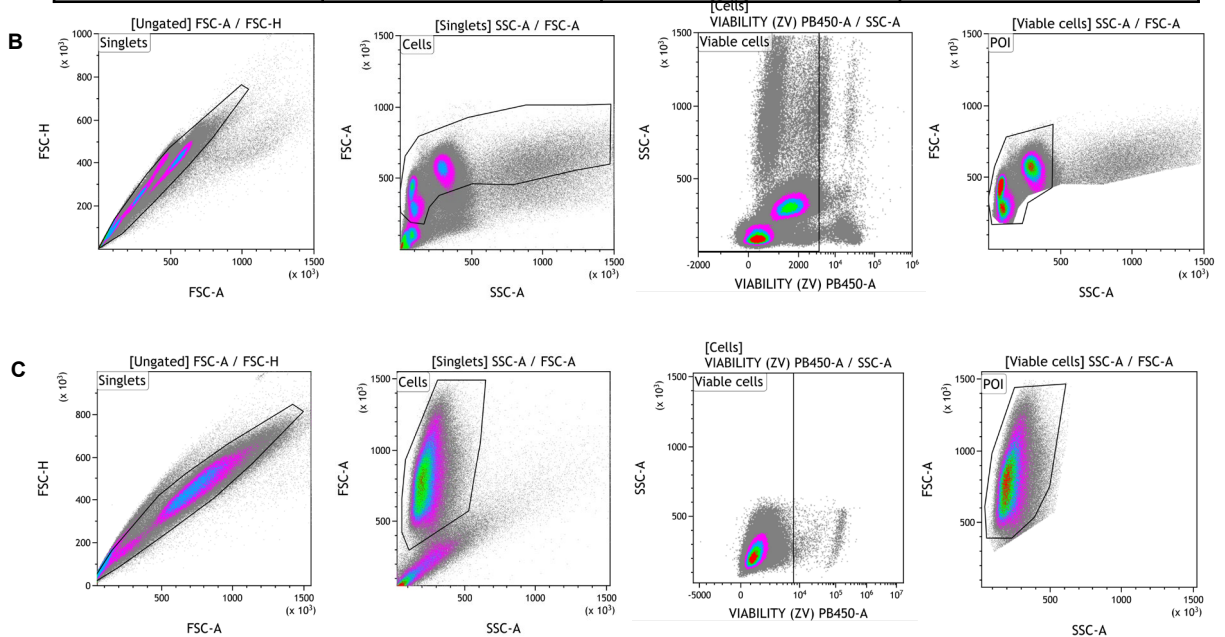
Figure S10: Volcano plot showing genes that were significantly up- (red dots, 108 genes) and downregulated (blue dots, 163 genes) between HEU and HUU samples based on their p-value (y-axis) and fold change (x-axis).

Figure S11: Full gene list displaying all 271 differentially expressed genes, fold change, p-value, gene symbol, and description.

Figure S12: Functional analysis of DEGs by Gene Ontology biological processes (GOBP) depicting 11 biological processes which are involved. Only upregulated but not downregulated genes were involved in these processes

Figure S1: Preliminary immunophenotypic analysis prior to Cytobank export and patient cohort details

	Sample volume (ml)	Purity after Run 1 (%)	Purity after Run 2 (%)
A			
HUU Sample 1	98	79,37	97,04
HUU Sample 2	75	72,08	98,29
HUU Sample 3	57	77,3	95,87
HUU Sample 4	108	21,98	98,65
HUU Sample 5	74	80,38	99,38
HEU Sample 1	64	66,95	96,64
HEU Sample 2	72	71,46	98,64
HEU Sample 3	67	60,89	95,02
HEU Sample 4	63	57,78	97,54



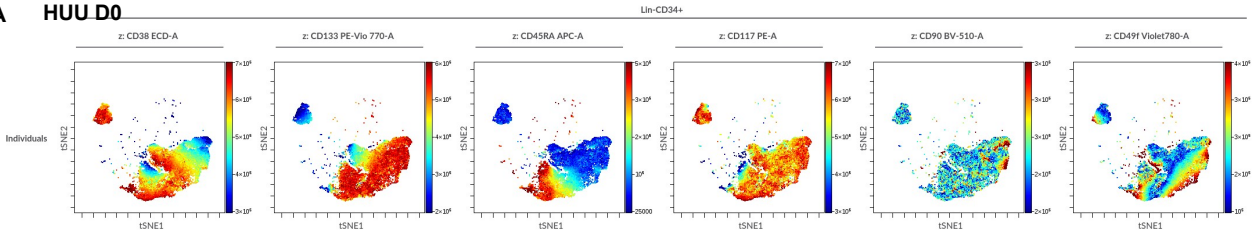
D

Sample	Maternal VL timepoint relative to delivery	VL result (copies/ml)	ARVs	HIV PCR in infant (timepoint relative to delivery)	HIV PCR result	Gene Xpert result on UCB
1	D+1 day	39	TLD	D	Negative	Negative
2	D-6 weeks	LTDL	TLD	D+72 hours	Negative	Negative
3	D	LTDL	TLD	D+96 hours	Negative	Negative
4	D-3 months	LTDL	TEE	D+1 day	Negative	Negative
5	D+1 day	79	TLD	D+1 day	Negative	Negative

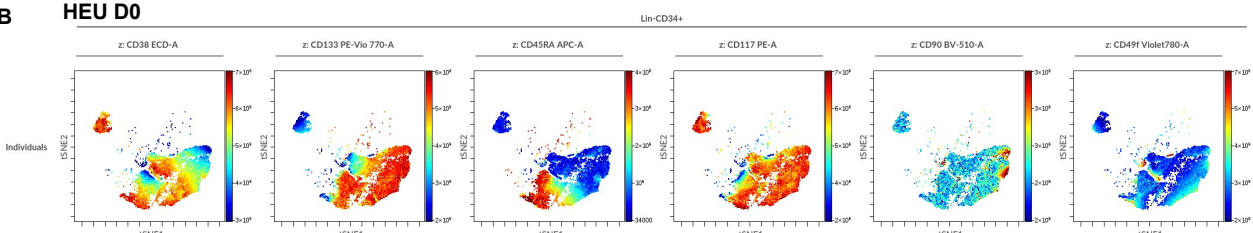
VL, Viral load; ARV-Antiretrovirals; D-Day of delivery; LTDL-Lower than detectable limit; PCR Polymerase chain reaction; TLD- TDF, Lamivudine, Dolutegravir; TEE- ;Tenofovir disoproxil fumarate(TDF), Emtricitabine and Efavirenz

Figure S2: Concatenated viSNE plots showing differences between HEU and HUU CD34+Lin- subsets

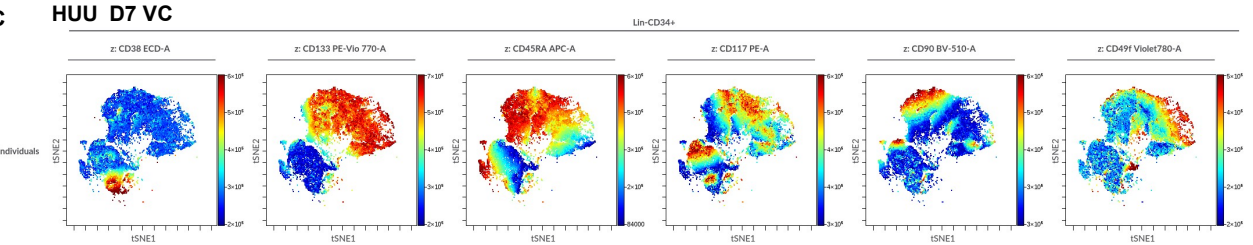
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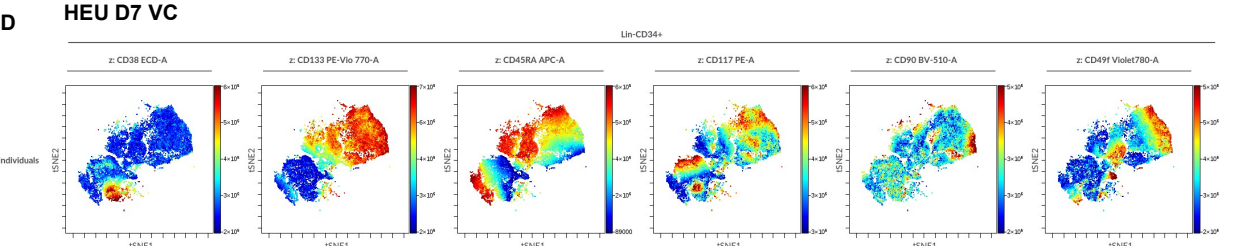
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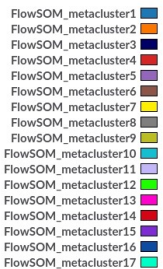
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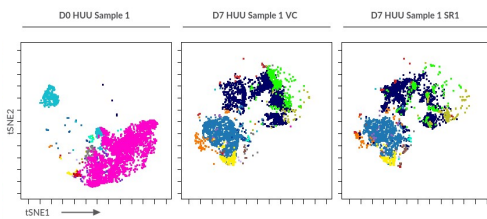
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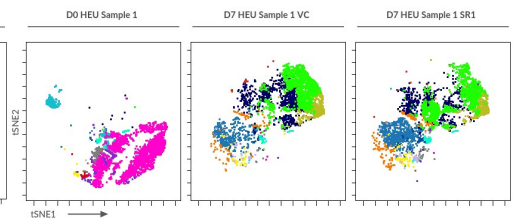
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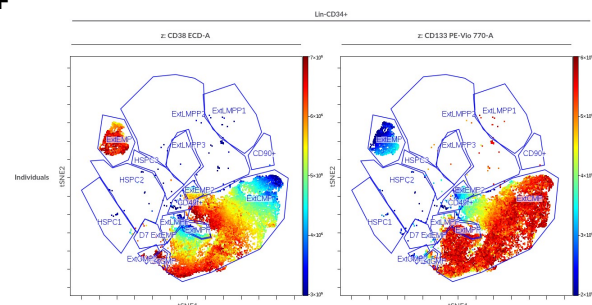
HUU



HEU



F



G

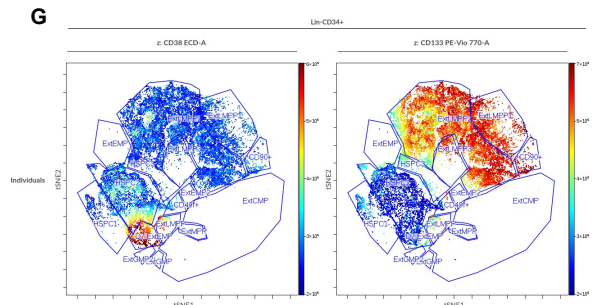


Fig. S3 Statistically significant metacluster differences in Lin-CD34+ sub-population between HEU and HUU populations.

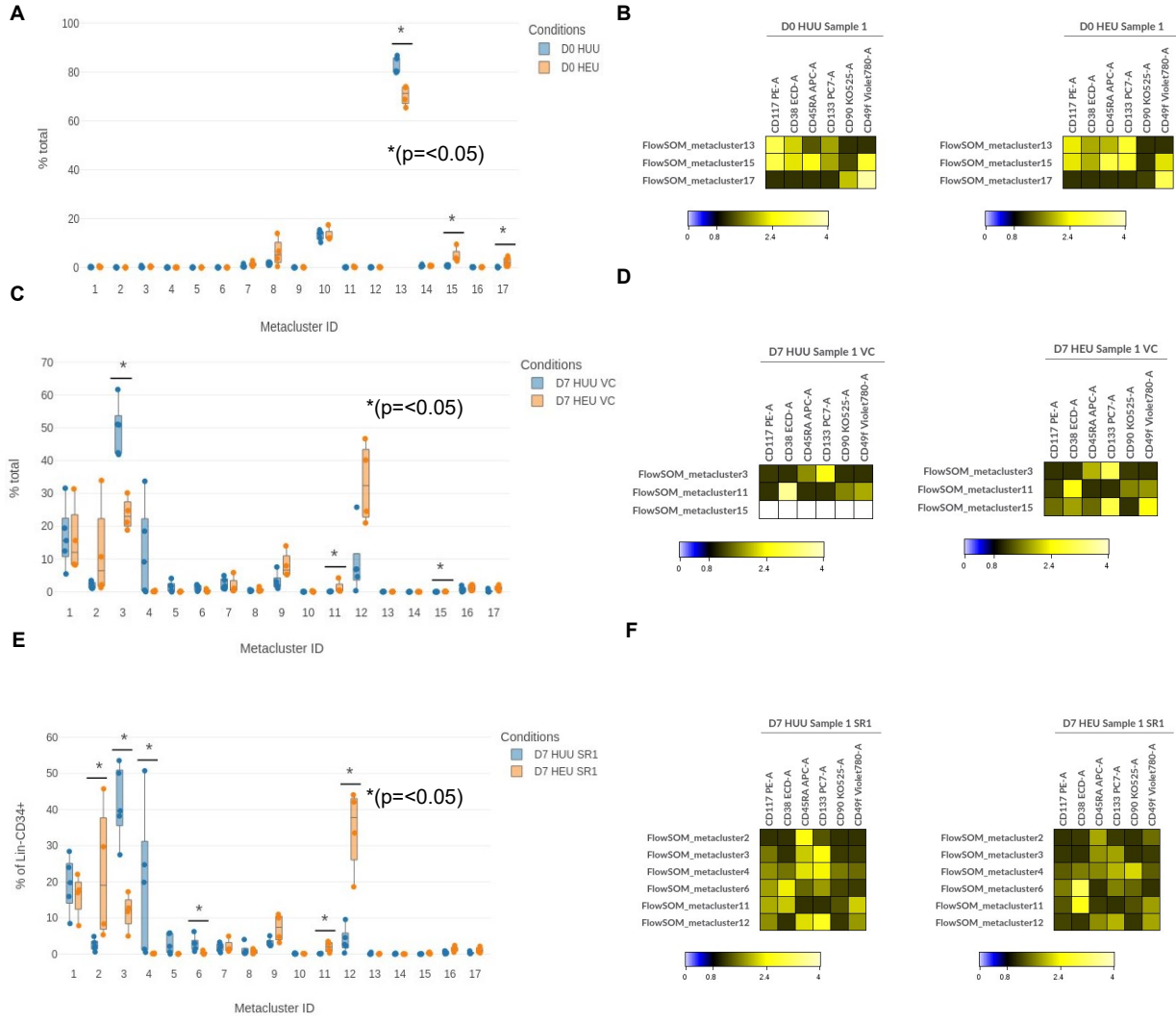
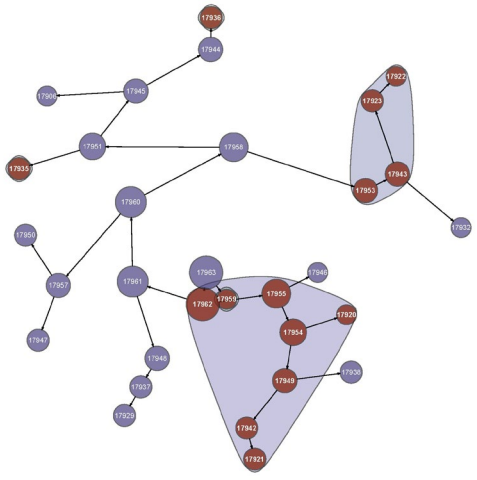
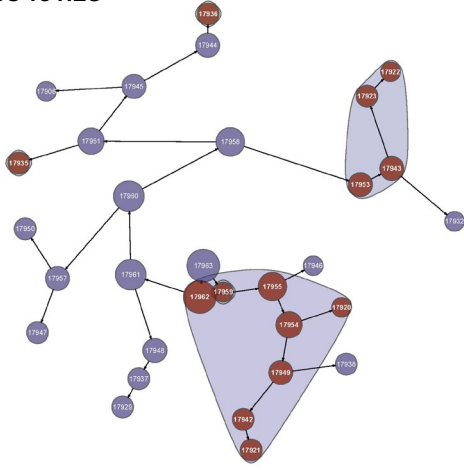
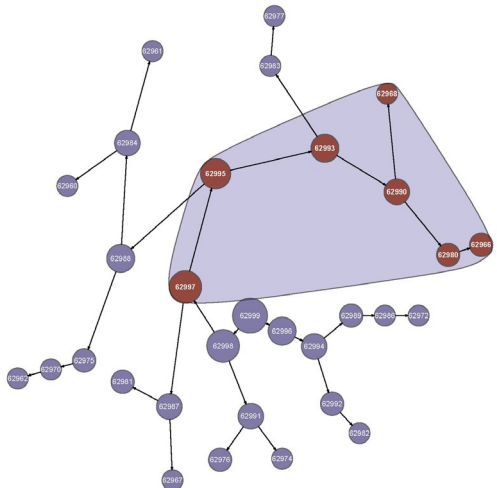
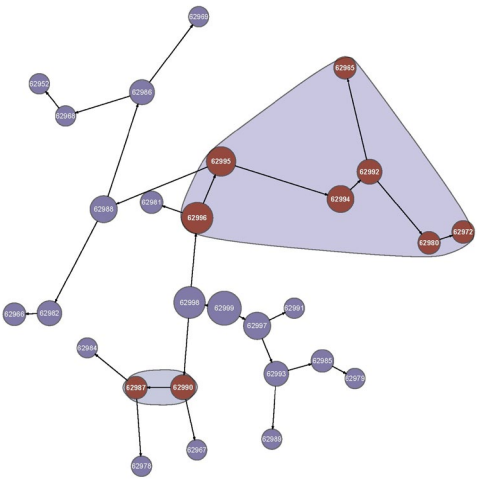


Figure S4: CITRUS cluster analysis showing differences between D0 and D7 HUU and HEU CD34+Lin- subsets

A D0 HUU vs HEU



B D7 VC HUU vs HEU



C D7 SR1 HUU vs HEU

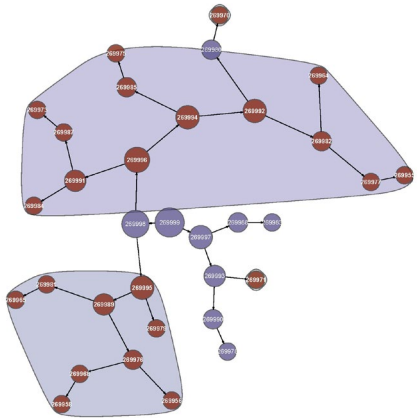
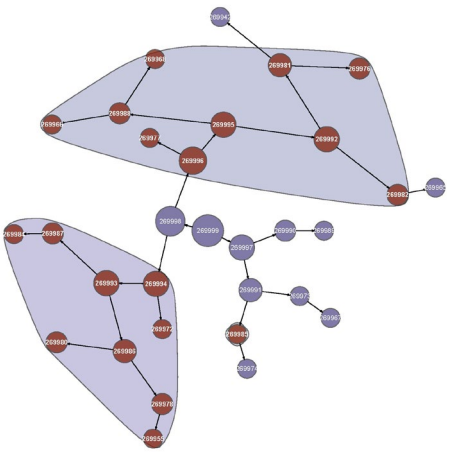
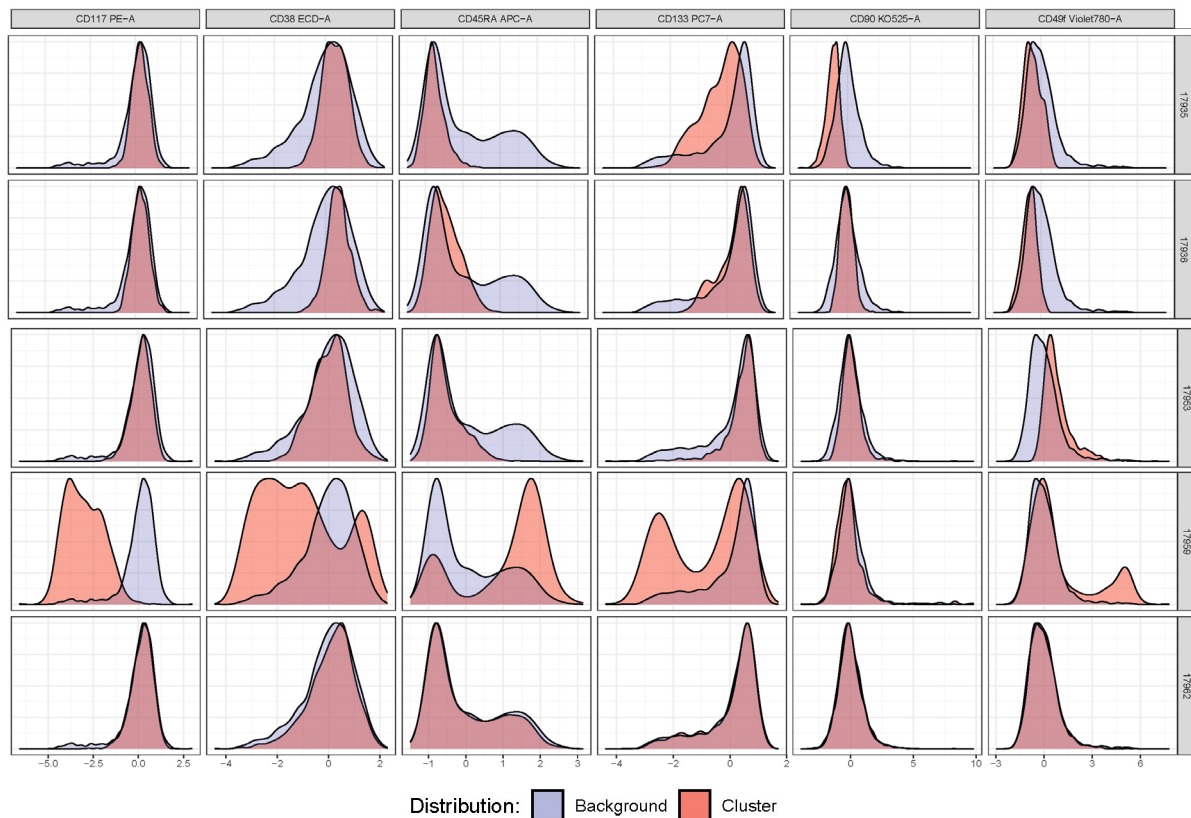


Fig. S5 D0 CITRUS cluster immunophenotype analysis of Lin-CD34+ sub-population between HEU and HUU subgroups.

A



B

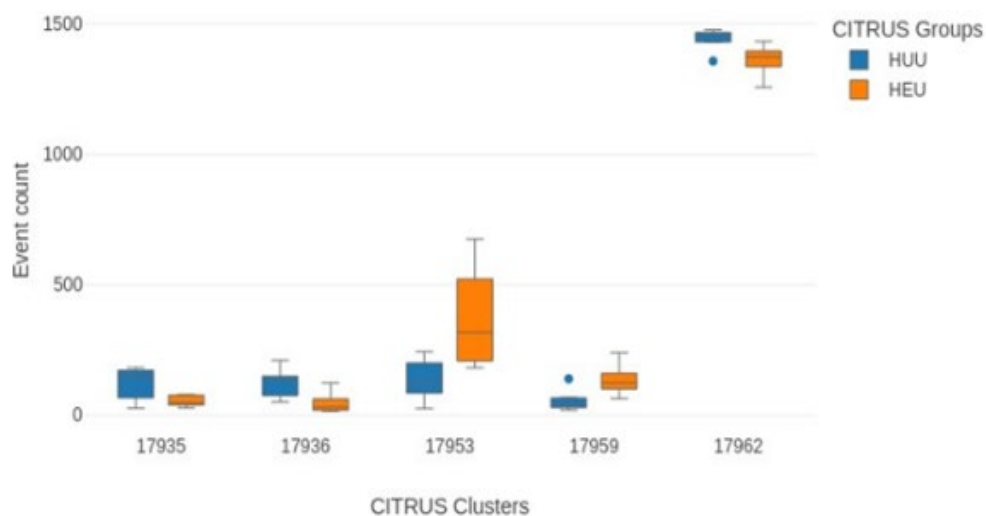
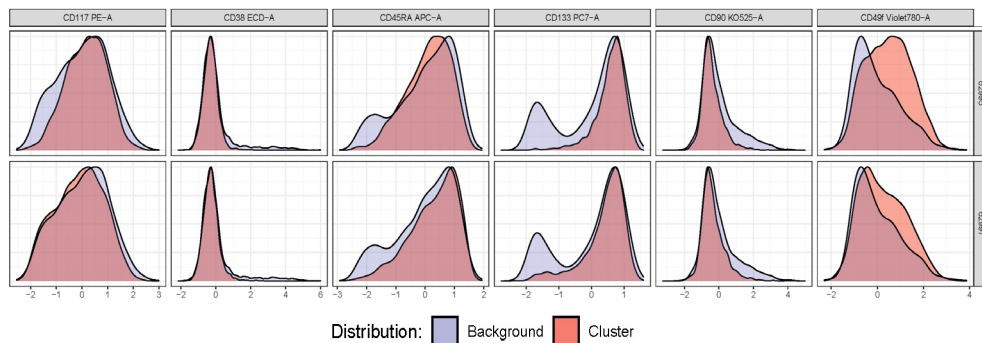
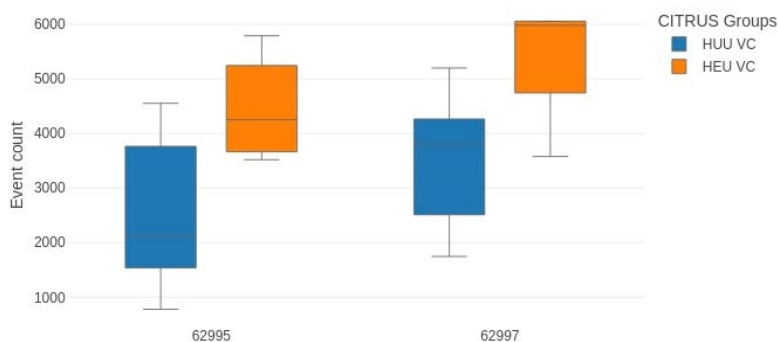


Fig. S6 D7 CITRUS cluster immunophenotype analysis of Lin-CD34+ sub-population between HEU and HUU subgroups.

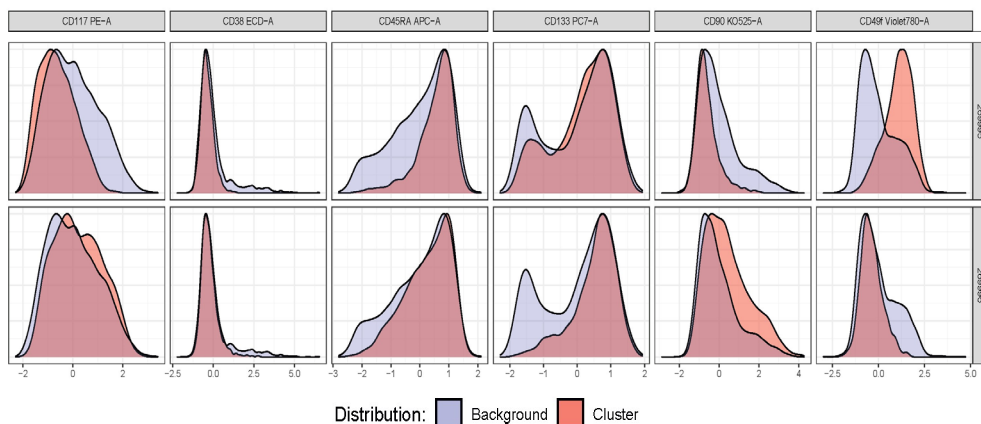
A



B



C



D

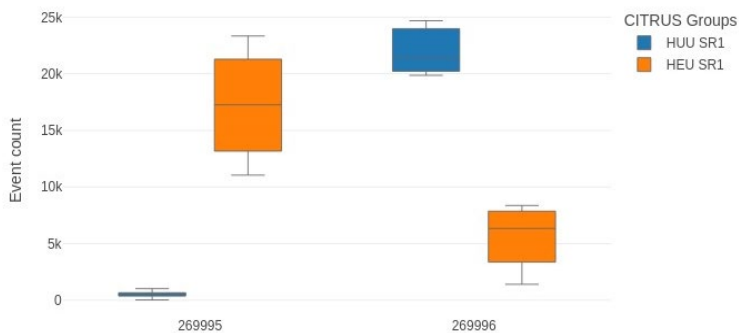


Figure S7: Concatenated UMAP plots showing differences between D0 and D7 HEU and HUU CD34-Lin- subsets

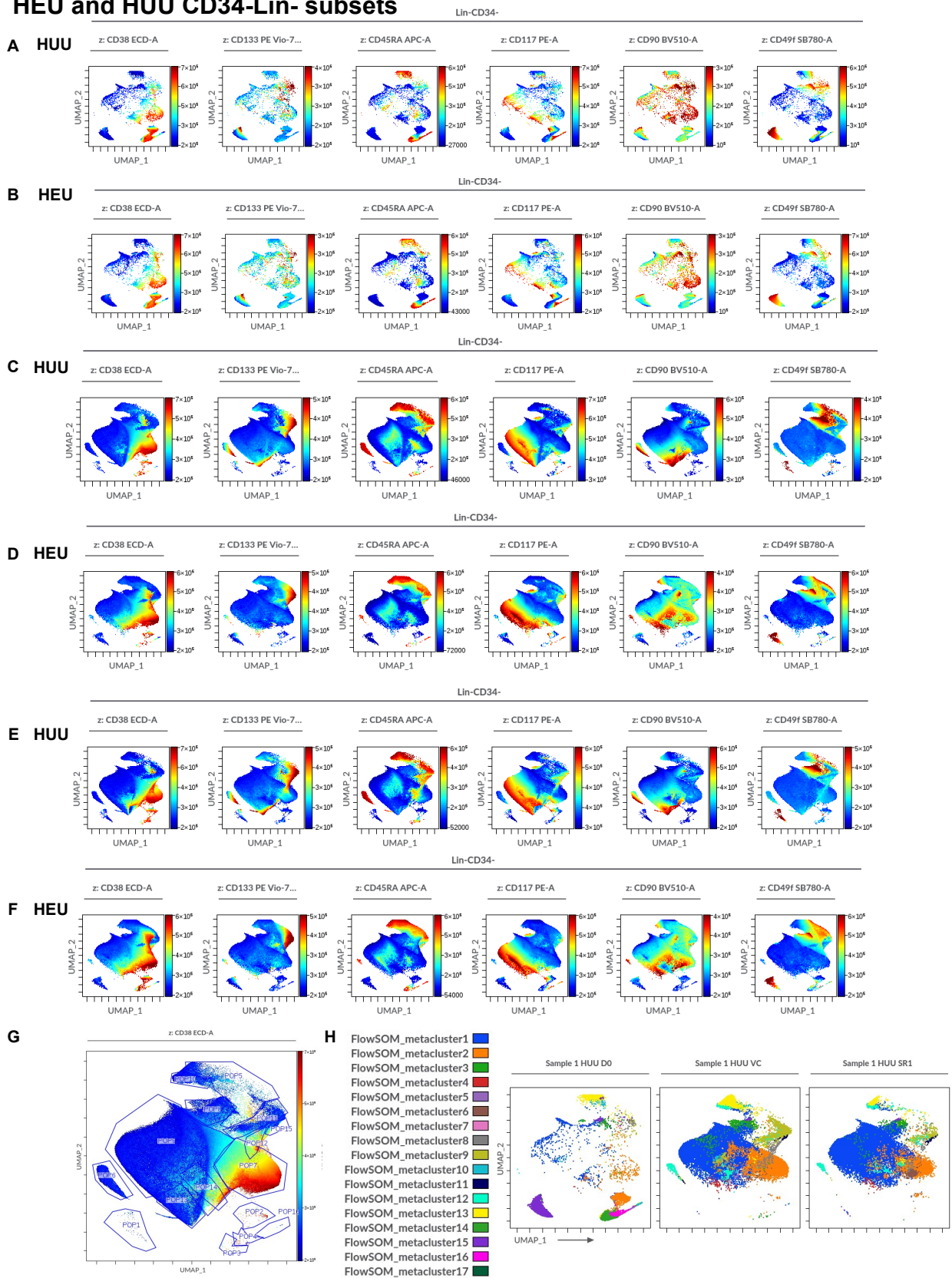


Figure S8: Population frequencies and statistically significant differences in Lin-CD34- sub-populations

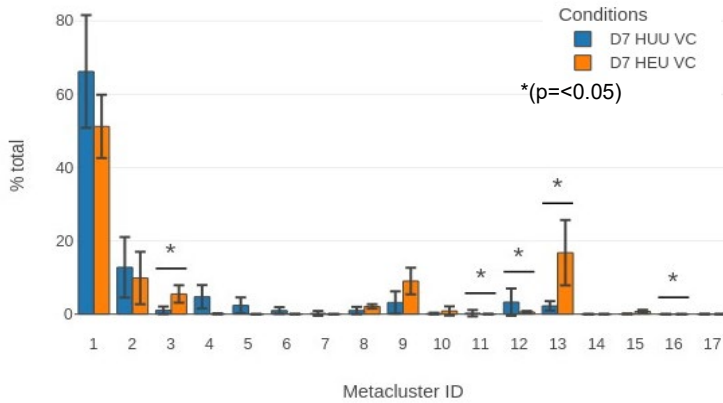
A

Average percentage of immunophenotypic sub-populations present in Lin-CD34- on D0 and D7 between HUU and HEU samples

Population name	Phenotype	% D0 HUU	% D0 HEU	% D7 VC HUU	% D7 VC HEU	% D7 SR1 HUU	% D7 SR1 HEU
POP1	Lin-CD34-CD38-CD133vCD45RA-CD117-CD90dim CD49f+	30,49	41,4	0,02	0,24	0,1	0,62
POP2	Lin-CD34-CD38+CD133dimCD45RA-CD117vCD90dimCD49f-	26,73	16,03	0,04	0,05	0,15	0,29
POP3	Lin-CD34-CD38vCD133-CD45RA+CD117-CD90-CD49f-	15,89	14,79	0,01	0,01	0,01	0,06
POP4	Lin-CD34-CD38+CD133-CD45RA+dimCD117-CD90dimCD49f-	11,04	11,35	0,01	0,01	0,05	0,04
POP5	Lin-CD34-CD38-CD133vCD45RA+CD117-CD90vCD49f+	3,41	5,47	2,53	20,29	2,12	38,41
POP6	Lin-CD34-CD38-CD133vCD45RA+CD117-CD90+ CD49f-	0	0	3,02	0,03	2,91	0,03
POP7	Lin-CD34-CD38+CD133-CD45RAvCD117vCD90vCD49f-	1,14	1,07	19,35	11,99	21,28	7,69
POP8	Lin-CD34-CD38-CD133-CD45RA-CD117vCD90dimCD49f+	0,65	0,74	4,14	9,94	3,24	7,45
POP9	Lin-CD34-CD38-CD133-CD45RAvCD117vCD90vCD49f-	4,37	4,65	56,03	41,68	53,64	31,49
POP10	Lin-CD34-CD38-CD133dimCD45RA+CD117-CD90+CD49f-	0,02	0,03	6,4	2,09	1,7	1,1
POP11	Lin-CD34-CD38vCD133vCD45RAvCD117vCD90vCD49fdim	0,23	0,15	2,61	7,32	1,75	7,06
POP12	Lin-CD34-CD38vCD133-CD45RA-CD117-CD90dimCD49fdim	0,12	0,08	1,24	1,71	1,6	2,02
POP13	Lin-CD34-CD38-CD133vCD45RA-CD117+CD90+CD49f-	0,01	0	7,26	0,28	11,89	0,49
POP14	Lin-CD34-CD38vCD133vCD45RA-CD117dimCD90+CD49f-	0,77	0,56	3,69	1,03	5,92	1,14
POP15	Lin-CD34-CD38-CD133+CD45RA+CD117+CD90dimCD49fdim	0	0	0,56	0,33	0,3	0,09
POP16	Lin-CD34-CD38dimCD133-CD45RA+CD117+CD90-CD49fv	3,27	2,28	0,01	0,03	0	0,07

CD, cluster of differentiation; CMP, common myeloid progenitor; D0, day 0; D7, day 7; EMP, erythro-myeloid progenitor; Ext, extended; GMP, granulocyte-macrophage progenitor; HSC, haematopoietic stem cell; LMPP, lympho-myeloid primed progenitor; MPP, multipotent progenitor; SR1, stemregenin1; v, variable; VC, vehicle control

B



C

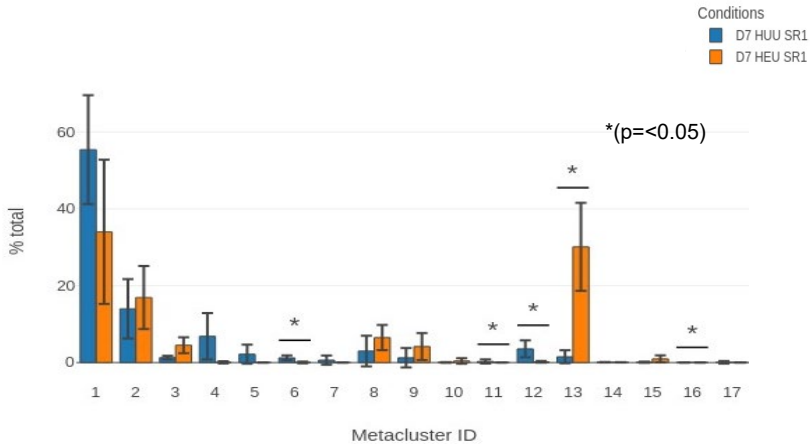
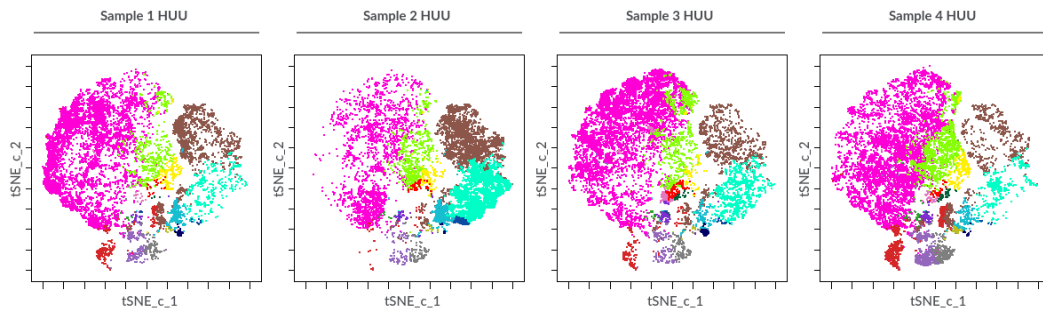


Figure S9: Differences in CFU phenotype between HEU and HUU samples.

A

HUU

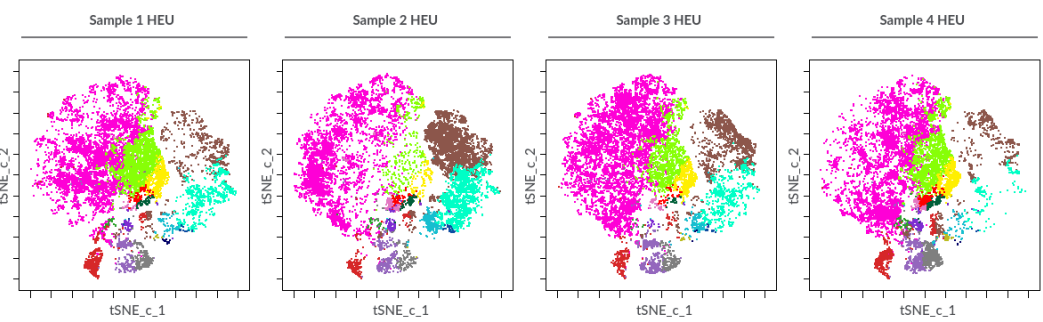
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- FlowSOM_metacluster2 ■
- FlowSOM_metacluster3 ■
- FlowSOM_metacluster4 ■
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- FlowSOM_metacluster7 ■
- FlowSOM_metacluster8 ■
- FlowSOM_metacluster9 ■
- FlowSOM_metacluster10 ■
- FlowSOM_metacluster11 ■
- FlowSOM_metacluster12 ■
- FlowSOM_metacluster13 ■
- FlowSOM_metacluster14 ■
- FlowSOM_metacluster15 ■
- FlowSOM_metacluster16 ■
- FlowSOM_metacluster17 ■



B

HEU

- FlowSOM_metacluster1 ■
- FlowSOM_metacluster2 ■
- FlowSOM_metacluster3 ■
- FlowSOM_metacluster4 ■
- FlowSOM_metacluster5 ■
- FlowSOM_metacluster6 ■
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- FlowSOM_metacluster8 ■
- FlowSOM_metacluster9 ■
- FlowSOM_metacluster10 ■
- FlowSOM_metacluster11 ■
- FlowSOM_metacluster12 ■
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- FlowSOM_metacluster15 ■
- FlowSOM_metacluster16 ■
- FlowSOM_metacluster17 ■



C

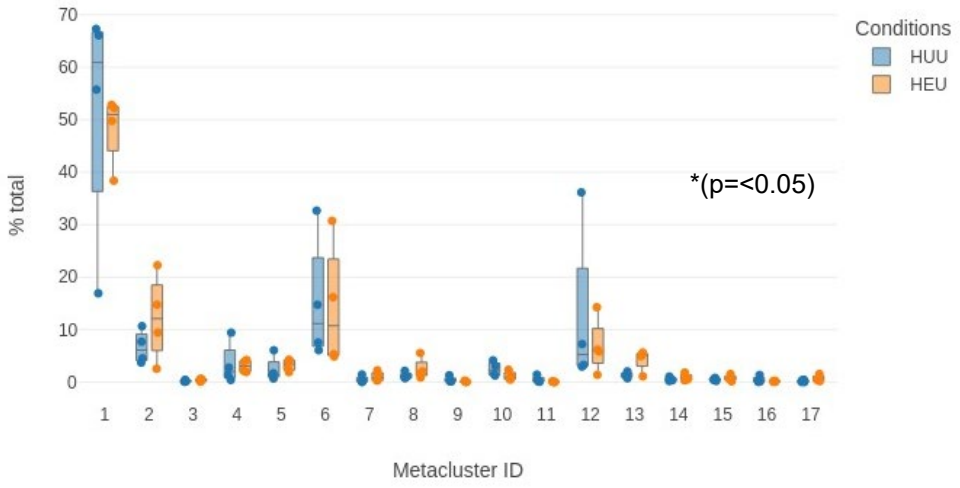


Figure S10: Volcano plot showing genes that were significantly up- (red dots, 108 genes) and downregulated (blue dots, 163 genes) between HEU and HUU samples based on their p-value (y-axis) and fold change (x-axis).

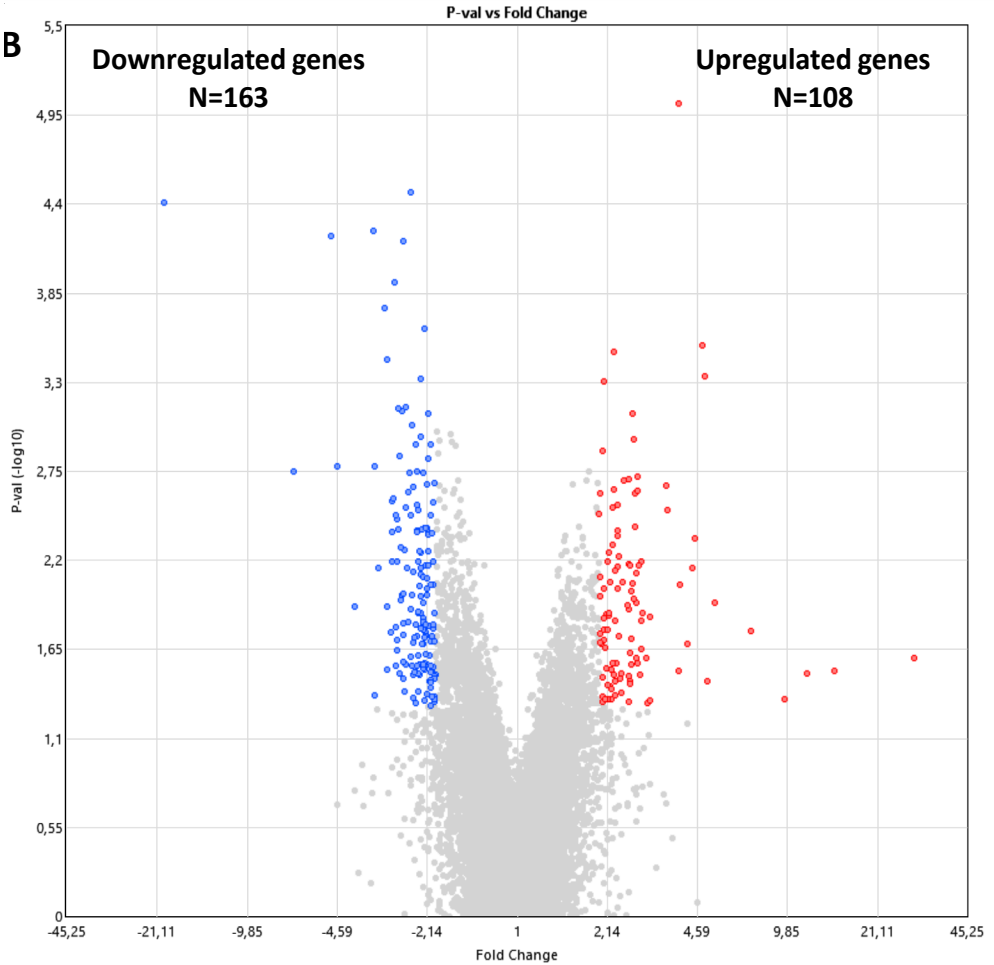


Figure S11: Full gene list displaying all 271 differentially expressed genes, fold change, p-value, gene symbol, and description.

ID	Fold Chang	P-val	Gene Symbol	Description
TC0100015858.hg.1	28.72	0,0255	S100A8	S100 calcium binding protein A8
TC0100015856.hg.1	14,66	0,0306	S100A12	S100 calcium binding protein A12
TC040007857.hg.1	11,66	0,0316	AREG	amphiregulin
TC1200008275.hg.1	9,64	0,0454	SYT1	synaptotagmin I
TC0X00010473.hg.1	7,24	0,0172	TSC22D8	TSC22 domain family, member 3
TC0200014414.hg.1	5,29	0,0114	CXCR4	chemokine (C-X-C motif) receptor 4
TC0200012429.hg.1	4,98	0,0253	PP1B	Memczak2013 ANTISENSE, coding, INTERNAL, intronic best transcript NM_0010393556
TC1200009800.hg.1	4,86	0,0005	SLC2A3	solute carrier family 2 (facilitated glucose transporter), member 3
TC1200009796.hg.1	4,78	0,0009	SLC2A14	solute carrier family 2 (facilitated glucose transporter), member 14
TC0100019392.hg.1	4,49	0,0046	S100A9	S100 calcium binding protein A9
TC1500007955.hg.1	4,42	0,007	AQP9	aquaporin 9
TC1200007652.hg.1	4,23	0,0205	GRASP	GRP1 (general receptor for phosphoinositides 1)-associated scaffold protein
TC1100006749.hg.1	3,95	0,009	ZNF215	zinc finger protein 215
TC0100015858.hg.1	3,91	5,55E-06	TXNIP	thioredoxin interacting protein
TSUnmapped00003954.hg.1	3,9	0,0032	EIF3F	Eukaryotic translation initiation factor 3 subunit F [Source:UniProtKB/Swiss-Prot;Acc:O03003]
TC0200015885.hg.1	3,56	0,0091	FAM124B	family with sequence similarity 124 member B
TC1200010006.hg.1	3,53	0,0022	PLBD1	phospholipase B domain containing 1
TC200009023.hg.1	3,08	0,0141	SAMHD1	SAM domain and HD domain 1
TC1900008900.hg.1	3,07	0,046	PPP6R1	Memczak2013 ANTISENSE, CDS, coding, INTERNAL best transcript NM_014591
TC1300008706.hg.1	3,01	0,0476	ELF1	Transcript identified by AceView, Entrez Gene ID(s) 1997; 100128628
TC0800011150.hg.1	2,97	0,0253	TP53NP1	tumor protein p53 inducible nuclear protein 1
TC1700007777.hg.1	2,9	0,0132	THRA	thyroid hormone receptor, alpha
TC1900011615.hg.1	2,86	0,0148	MRAR7	matrix-remodelling associated 7
TC2200008661.hg.1	2,82	0,0222	LALSL2	lectin, galactoside-binding, soluble, 2
TC0100010241.hg.1	2,85	0,0063	MINDA	myeloid cell nuclear differentiation antigen
TC1700012262.hg.1	2,81	0,0323	PSME3	proteasome activator subunit 3
TC2200008475.hg.1	2,79	0,0067	PIK3P1	phosphoinositide-3-kinase interacting protein 1
TC0700013442.hg.1	2,78	0,0019	LSMEM1	leucine-rich single-pass membrane protein 1
TC0700011584.hg.1	2,78	0,0273	FGL2	fibrinogen-like 2
TC1900008881.hg.1	2,76	0,0023	FCAR	Fc fragment of IgA receptor
TC0800009819.hg.1	2,75	0,0115	DOK2	docking protein 2
TC0300010770.hg.1	2,75	0,025	CSRNP1	cysteine-serine-rich nuclear protein 1
TC1200010559.hg.1	2,73	0,0076	VDR	vitamin D (1,25- dihydroxyvitamin D3) receptor
TC0600013231.hg.1	2,7	0,0024	SGK1	serum/glucocorticoid induced kinase 1
TC2000007117.hg.1	2,7	0,0039	ASXL1	additional sex combs like transcriptional regulator 1
TC1900008141.hg.1	2,68	0,0002	CYP251	cytochrome P450, family 2, subfamily S, polypeptide 1
TC0100016944.hg.1	2,68	0,0011	ARL8A	ADP-ribosylation factor like GTPase 8A
TC0400078513.hg.1	2,65	0,0068	EPGN	epithelial mitogen
TC1700011910.hg.1	2,65	0,0008	CYTH1	cytohesin 1
TC0500011852.hg.1	2,64	0,0276	LOX	lysyl oxidase
TC0700010604.hg.1	2,64	0,0279	CPVL	carboxypeptidase, vitellogenic-like
TC0200010795.hg.1	2,63	0,0097	SLC11A1	solute carrier family 11 (proton-coupled divalent metal ion transporter), member 1
TC2200007356.hg.1	2,61	0,0193	KDELR3	KDEL (Lys-Asp-Glu-Leu) endoplasmic reticulum protein retention receptor 3
TC0500012238.hg.1	2,61	0,0234	HBEGF	heparin-binding EGF-like growth factor
TC0300007016.hg.1	2,6	0,0351	ARPP21	Transcript identified by AceView, Entrez Gene ID(s) 10777
TC1500010160.hg.1	2,59	0,0068	CTSH	cathepsin H
TC0400007853.hg.1	2,59	0,0365	EREG	epiregulin
TC0100008625.hg.1	2,58	0,0127	PDE4B	phosphodiesterase 4B, cAMP-specific
TC1100013012.hg.1	2,58	0,002	STX3	syntaxin 3
TC1500010731.hg.1	2,58	0,047	TRAF69	tripartite motif containing 69
TC0500007138.hg.1	2,58	0,0326	IL7R	interleukin 7 receptor
TC0600012009.hg.1	2,58	0,0067	DEFB114	defensin, beta 114
TC0500012519.hg.1	2,55	0,0118	SPARC	secreted protein, acidic, cysteine-rich (osteonectin)
TC0100015789.hg.1	2,47	0,002	POGZ	Transcript identified by AceView, Entrez Gene ID(s) 23126
TC0700013581.hg.1	2,43	0,0086	STEAP4	STEAP family member 4
TC0M00006437.hg.1	2,42	0,0313	COX2	cytochrome c oxidase subunit II
TC0600008109.hg.1	2,4	0,0415	VEGFA	vascular endothelial growth factor A
TC1800008743.hg.1	2,4	0,0396	CCDC68	coiled-coil domain containing 68
TC0600011843.hg.1	2,36	0,0185	TBOC	tubulin folding cofactor C
TC1700009528.hg.1	2,36	0,006	CXCL16	chemokine (C-X-C motif) ligand 16
TC0200009662.hg.1	2,34	0,0029	TNFAIP6	tumor necrosis factor, alpha-induced protein 6
TC0300008550.hg.1	2,34	0,0068	CSTA	cystatin A (stefin A)
TC0300011516.hg.1	2,34	0,0045	EIF4E9	eukaryotic translation initiation factor 4E family member 3
TC0800007284.hg.1	2,33	0,0094	KCNU1	potassium channel, subfamily U, member 1
TC0200007261.hg.1	2,32	0,0041	QPCT	glutamyl-peptide cyclotransferase
TC1000009865.hg.1	2,31	0,027	FAM107B	Memczak2013 ALT_ACCEPTOR, ALT_DONOR, coding, INTERNAL, intronic best transcript NM_031453
TC0700011571.hg.1	2,29	0,0073	YWHAG	Memczak2013 ALT_ACCEPTOR, ALT_DONOR, coding, INTERNAL, intronic best transcript NM_012479
TC0300009692.hg.1	2,28	0,0148	DVL3	dishevelled segment polarity protein 3
TC1000011807.hg.1	2,28	0,0433	SMNDC1	survival motor neuron domain containing 1
TC1200011961.hg.1	2,28	0,0349	TMEM116	transmembrane protein 116
TC0100007832.hg.1	2,26	0,0003	C3H12A; MIR6732	zinc finger CCHC-type containing 12A; microRNA 6732
TC1700011523.hg.1	2,26	0,0023	PSMD12	proteasome 26S subunit, non-ATPase 12
TC0500010265.hg.1	2,26	0,0318	CDH18	cadherin 18, type 2
TC1100010809.hg.1	2,24	0,003	ORAC12	olfactory receptor, family 4, subfamily C, member 12
TC0600013273.hg.1	2,23	0,0051	BCLAF1	Memczak2013 ALT_DONOR, coding, INTERNAL, intronic best transcript NM_014739
TC1400010085.hg.1	2,23	0,0273	TCL1A	T-cell leukemia/lymphoma 1A
TC0300012016.hg.1	2,23	0,0455	P2281	pumilio-like protein 228, 5 protein coupled, 12
TC0400012178.hg.1	2,23	0,0297	DCHS2	dachsous cadherin-related 2
TC0600012562.hg.1	2,21	0,0395	BACH2	Memczak2013 ALT_ACCEPTOR, ALT_DONOR, coding, INTERNAL, intronic best transcript NM_0011070794
TC0600014360.hg.1	2,2	0,0085	IPCEF1	interaction protein for cytohesin exchange factors 1
TC0200007200.hg.1	2,18	0,0375	LTBP1	latent transforming growth factor beta binding protein 1
TC1100006831.hg.1	2,18	0,0132	ADM	adrenomedullin
TC0600013570.hg.1	2,18	0,0056	ZBTB2	zinc finger and BTB domain containing 2
TC0900008148.hg.1	2,17	0,0138	TDRD7	tudor domain containing 7
TC1600007936.hg.1	2,14	0,0065	SLCGA2	solute carrier family 6 (neurotransmitter transporter), member 2
TC1500006811.hg.1	2,14	0,037	CHRM5	cholinergic receptor, muscarinic 5
TC1100010733.hg.1	2,14	0,0458	SP1	Sp1 proto-oncogene
TC1200007950.hg.1	2,14	0,0168	IAPP	islet amyloid polypeptide
TC0700012467.hg.1	2,13	0,0136	TMEM229A	transmembrane protein 229A
TC0700008190.hg.1	2,12	0,0292	CD36	CD36 molecule (thrombospondin receptor)
TC1600011426.hg.1	2,11	0,0452	ATRXN1L	ataxin 1-like
TC0500010553.hg.1	2,09	0,0217	FYB	FYB binding protein
TC0100008026.hg.1	2,09	0,0169	YBX1	Zhang2013 ALT_ACCEPTOR, ALT_DONOR, coding, INTERNAL, intronic best transcript NM_004559
TC1000007883.hg.1	2,09	0,0195	SRGN	serglycin
TC0100006805.hg.1	2,08	0,0094	RBP7	retinol binding protein 7, cellular

TC1900010625.hg.1	2,08	0,0143 LGL54	lectin, galactoside-binding, soluble, 4
TC1700007415.hg.1	2,07	0,0005 CRYBA1	crystallin beta A1
TC1300006676.hg.1	2,06	0,0013 RASL11A	RAS4-like, family 11, member A
TC1600006446.hg.1	2,05	0,0434 HBA1	hemoglobin, alpha 1
TC0500012486.hg.1	2,05	0,0473 DCTN4	dynactin 4 [p82]
TC1100009904.hg.1	2,05	0,0381 OR51A2	olfactory receptor, family 51, subfamily A, member 2
TC1200012753.hg.1	2,04	0,0207 KLRC2	killer cell lectin-like receptor subfamily C, member 2
TC1200006561.hg.1	2,02	0,0024 RAD51AP1	RAD51 associated protein 1
TC1400007890.hg.1	2,01	0,0105 GPR65	G protein-coupled receptor 65
TC2100007821.hg.1	2,01	0,0201 ADAMTS1	ADAM metalloproteinase with thrombospondin type 1 motif 1
TC0400002649.hg.1	2,01	0,008 N56	NADH dehydrogenase, subunit 6 (complex I)
TC2000008666.hg.1	2,01	0,018 NAR8	N-ethylmaleimide-sensitive factor attachment protein, beta
TC1800007063.hg.1	2	0,0039 DTNA	Transcript identified by AceView, Entrez Gene ID(s) 1837
TC0100017722.hg.1	-2	0,0321 C10orf131	chromosome 1 open reading frame 131
TSUnmapped0000306.hg.1	-2,01	0,02 ZKSCAN7	zinc finger with KRAB and SCAN domains 7
TC0600009143.hg.1	-2,01	0,0336 SLC16A10	solute carrier family 16 (aromatic amino acid transporter), member 10
TC0500009010.hg.1	-2,01	0,0327 TERC61	Transcript identified by AceView, Entrez Gene ID(s) 10915
TC0100008631.hg.1	-2,01	0,0473 SGIP1	SH3-domain GRB2-like (endophilin) interacting protein 1
TC0200016659.hg.1	-2,02	0,0451 ATP6V1E2	ATPase, H+ transporting, lysosomal 31kDa, V1 subunit E2
TC1200006557.hg.1	-2,02	0,0138 TIGAR	TP53 induced glycolysis regulatory phosphatase
TC0200007195.hg.1	-2,02	0,0434 TTC27; MIR4765	tetratricopeptide repeat domain 27; microRNA 4765
TC0500013360.hg.1	-2,03	0,0021 CDK13	cyclin-dependent kinase-like 3
TC1400010737.hg.1	-2,03	0,0028 KLHL28	kelch-like family member 28
TC0300012339.hg.1	-2,04	0,0089 GATA2	GATA binding protein 2
TC0400012261.hg.1	-2,04	0,0064 NUDT16	nucleic acid hydrolase 6
TC0400012496.hg.1	-2,04	0,0205 AGA	aspartylglucosaminidase
TC0800010490.hg.1	-2,04	0,0165 TMEM68	transmembrane protein 68
TC1100012187.hg.1	-2,04	0,0158 KBTBD9	kelch repeat and BTB (POZ) domain containing 3
TC1700012231.hg.1	-2,05	0,0435 TMEM98	transmembrane protein 98
TC0100011421.hg.1	-2,05	0,0298 CD46	Transcript identified by AceView, Entrez Gene ID(s) 4179
TC1900011922.hg.1	-2,06	0,0185 ZNF728	zinc finger protein 728
TC1600009514.hg.1	-2,07	0,0042 RRN3	RRN3 homolog, RNA polymerase I transcription factor
TC0100009035.hg.1	-2,07	0,0012 KIAA1107	KIAA1107
TC1200012610.hg.1	-2,07	0,0346 SSPN	sarcospan
TC0600013574.hg.1	-2,07	0,0039 RMN1	required for meiotic nuclear division 1 homolog
TC1900007267.hg.1	-2,07	0,009 AP1M1	adaptor-related protein complex 1, mu 1 subunit
TC0700002680.hg.1	-2,07	0,0494 TMEM176A	transmembrane protein 176A
TC1200012706.hg.1	-2,07	0,0307 ACAD10	acyl-CoA dehydrogenase family, member 10
TC1300007406.hg.1	-2,07	0,0281 IRAK4	interleukin 1 receptor associated kinase 4
TC1700012283.hg.1	-2,07	0,0306 LUC7L3	LUC7-like 3 pre-mRNA splicing factor
TC1200007706.hg.1	-2,07	0,0199 MFS05	major facilitator superfamily domain containing 5
TC1900011754.hg.1	-2,08	0,0355 ZNF225	zinc finger protein 225
TC0500010628.hg.1	-2,09	0,0354 ANXA2R	annexin A2 receptor
TC0100018367.hg.1	-2,11	0,0242 CHR1M3	cholinergic receptor, muscarinic 3
TC1700009402.hg.1	-2,12	0,016 SMYD4	SET and MYND domain containing 4
TC1000009894.hg.1	-2,13	0,0044 FAM188A	family with sequence similarity 188, member A
TC1700008350.hg.1	-2,13	0,0067 NOG	noggin
TC1100007266.hg.1	-2,13	0,0015 PDHX	pyruvate dehydrogenase complex, component X
TC1900011936.hg.1	-2,13	0,0055 ZNF781	zinc finger protein 781
TC0200016884.hg.1	-2,14	0,0008 TIA1	TIA1 cytotoxic granule-associated RNA binding protein
TC0200010936.hg.1	-2,14	0,042 ZFAND2B	zinc finger, AN1-type domain 2B
TC0600008615.hg.1	-2,14	0,004 KNCU35	hes2013 ALT_ACCEPTOR, ALT_DONOR, coding, INTERNAL, intronic best transcript NM_001160133
TC1300009999.hg.1	-2,15	0,0061 BINM	basic, immunoglobulin-like variable motif containing
TC1900008300.hg.1	-2,15	0,0104 RELB	v-rel avian reticuloendotheliosis viral oncogene homolog B
TC0700012159.hg.1	-2,15	0,0094 PUS7	pseudouridylyl synthase 7 (putative)
TC0300009613.hg.1	-2,15	0,004 USP13	ubiquitin specific peptidase 13 (isopeptidase T-3)
TC0300006627.hg.1	-2,15	0,0021 TSEN2	TSEN2 tRNA splicing endonuclease subunit
TC0600014246.hg.1	-2,16	0,0277 C6orf229	chromosome 6 open reading frame 229
TC0300006781.hg.1	-2,17	0,0176 KCNH8	potassium channel, voltage-gated eag related subfamily H, member 8
TC1900012025.hg.1	-2,17	0,0157 ZNF28	zinc finger protein 28
TC0900010792.hg.1	-2,18	0,0068 NOL8	nucleolar protein 8
TC1100013000.hg.1	-2,18	0,0185 CLP1; AP000662.4	Transcript identified by AceView, Entrez Gene ID(s) 10978; novel transcript, sense overlapping CLP1
TC0300014077.hg.1	-2,18	0,0236 WDR49	WD repeat domain 49
TC1800008097.hg.1	-2,18	0,004 MRPPE1	metallophosphoesterase 1
TC0500011173.hg.1	-2,19	0,0461 POC5	POC5 centriolar protein
TC1900009752.hg.1	-2,19	0,0272 PRDX2	peroxiredoxin 2
TC1700009254.hg.1	-2,19	0,0189 HXDC6	hexosaminidase (glycosyl hydrolase family 20, catalytic domain) containing
TC0400006756.hg.1	-2,19	0,004 TRMT144	RNA methyltransferase 44 homolog (S. cerevisiae)
TC0700008094.hg.1	-2,2	0,0002 ZP3	zona pellucida glycoprotein 3 (sperm receptor)
TC0100011064.hg.1	-2,2	0,03 CFH	complement factor H
TC1300007731.hg.1	-2,21	0,008 GPR180	G protein-coupled receptor 180
TC1200008113.hg.1	-2,22	0,0152 NUP107	nucleoporin 107kDa
TC0600011517.hg.1	-2,22	0,0207 HLA-DPA1	major histocompatibility complex, class II, DP alpha 1
TC0900011523.hg.1	-2,22	0,0168 HSPA5	heat shock 70kDa protein 5 (glucose-regulated protein, 78kDa)
TC0800008235.hg.1	-2,22	0,0161 FAM92A1	family with sequence similarity 92, member A1
TC2000009324.hg.1	-2,22	0,0276 DRICH1	aspartate-rich 1
TC0400011749.hg.1	-2,22	0,0293 BBS5	Bardet-Biedl syndrome 7
TC2000008571.hg.1	-2,22	0,0114 LRSPL	LDL receptor related protein 5 like
TC0100016715.hg.1	-2,23	0,0145 PTGS2	prostaglandin-endoperoxidase synthase 2 (prostaglandin G/H synthase and cyclooxygenase)
TC0300013687.hg.1	-2,23	0,0018 TRFC	Transcript identified by AceView, Entrez Gene ID(s) 7037
TC0800010553.hg.1	-2,23	0,0207 NSAAF	neuronal sphingomyelinase activation associated factor
TC1400008362.hg.1	-2,23	0,004 TRAF3	TNF receptor-associated factor 3
TC0200010516.hg.1	-2,26	0,0134 CYP20A1	cytochrome P450, family 20, subfamily A, polypeptide 1
TC0700011601.hg.1	-2,26	0,0077 TMEM60	transmembrane protein 60
TC1000006863.hg.1	-2,26	0,0165 RPP38	ribonuclease P/MRP 38kDa subunit
TC0400009458.hg.1	-2,27	0,0056 CDKN2AIP	CDKN2A interacting protein

TC0100011746.hg.1	-2,27	0,0165 CN1H4	corrichon family AMPA receptor auxiliary protein 4
TC0900008804.hg.1	-2,27	0,0071 ZNF79	zinc finger protein 79
TC1900010873.hg.1	-2,27	0,0005 ZNF45	zinc finger protein 45
TC1200008843.hg.1	-2,27	0,0105 FAM216A	family with sequence similarity 216, member A
TC1200010641.hg.1	-2,27	0,0166 BCDIN3D	BCDIN3 domain containing
TC0400009503.hg.1	-2,28	0,0011 PRIMPOL	primase and DNA directed polymerase
TC0700008650.hg.1	-2,28	0,0056 FAM185A	family with sequence similarity 185, member A
TC0100118446.hg.1	-2,28	0,0275 OMA1	OMA1 zinc metallopeptidase
TC0400008947.hg.1	-2,3	0,032 TMEM184C	transmembrane protein 184C
TC1100008091.hg.1	-2,3	0,009 CC5	copper chaperone for superoxide dismutase
TC1400007265.hg.1	-2,3	0,0316 TMEM260	transmembrane protein 260
TC1200112636.hg.1	-2,3	0,0064 C12orf73	chromosome 12 open reading frame 73
TC1000007427.hg.1	-2,3	0,0405 ZNF485	zinc finger protein 485
TC0400012938.hg.1	-2,31	0,0132 RASGEF1B	RASGEF domain family member 1B
TC0200008923.hg.1	-2,31	0,0139 PLGLB2	plasminogen-like B2
TC1100011938.hg.1	-2,32	0,0245 MRGPRD	MAS-related GPR, member D
TC0900012384.hg.1	-2,32	0,0091 H1FX	H1 histone family, member X
			<p> Homo sapiens family with sequence similarity 103, member A1, mRNA (cDNA clone MGC:2560 IMAGE:2989772), complete cds; Homo sapiens family with sequence similarity 103, member A1, mRNA (cDNA clone MGC:18025 IMAGE:3924570), complete cds; Homo sapiens family with sequence similarity 103, member A1, mRNA (cDNA clone MGC:102778 IMAGE:5578103), complete cds. T cell activation inhibitor, mitochondrial Jek2/2013 ANTISENSE, coding, INTERNAL, intronic best transcript NM_003380 calpain 3 TRG2; TRGV9; TRG1C; TRG1P; TRGIP2 T cell receptor gamma joining 1; T cell receptor gamma constant 2; T cell receptor gamma joining 2; T cell receptor gamma variable 9; T cell receptor gamma constant 1; T cell receptor gamma joining 2; T cell receptor gamma joining P2 EF-hand calcium binding domain 7; deleted in lymphocytic leukemia 2-like 0,0028 EFCAB7; DLEU2L histone cluster 2, H2aa4; histone cluster 2, H2aa3 0,0012 TMEM1177 transmembrane protein 1177 0,0156 FGGY FGGY carbohydrate kinase domain containing 0,048 MVH3 myosin, heavy chain 3, skeletal muscle, embryonic 0,0309 ZNF528 zinc finger protein 528 0,0188 B4GALT5 UDP-GalbetaGlcNAc beta 1,4-galactosyltransferase, polypeptide 5 -2,4 0,0022 NUDT17 nudix hydrolase 17 -2,4 0,0074 CDAN1 codanin 1 -2,4 0,0449 CATSPERG catpser channel auxiliary subunit gamma -2,41 0,032 RRP12 Memczak2018 ALT_ACCEPTOR, ALT_DONOR, coding, INTERNAL, intronic best transcript NM_001145114 -2,42 0,0202 ALG13 Transcript identified by AceView, Entrez Gene ID(s) 79868 -2,43 0,0104 ARHGEF18 Rho/Rac guanine nucleotide exchange factor 18 -2,43 0,028 REPS2 RALBP1 associated Eps domain containing 2 -2,45 0,0009 CMTM8 CKLF-like MARVEL transmembrane domain containing 8 -2,46 0,0125 ZNF720 zinc finger protein 720 -2,46 0,0249 CTSE cathepsin F -2,46 3,32E-05 HIST3H3A histone cluster 1, H3a -2,47 0,0093 DDX59 DEAD (Asp-Glu-Ala-Asp) box polypeptide 59 -2,49 0,0018 FASTKD3 FAST kinase domains 3 -2,51 0,0024 NEURL2 neuronal E3 ubiquitin protein ligase 2 -2,53 0,0152 PTCO2 pentatricopeptide repeat domain 2 -2,54 0,007 H1FO H1 histone family, member D -2,56 0,0029 CODC77 coiled-coil domain containing 77 -2,57 0,0007 WRAP73 WD repeat containing, antisense to TP73 -2,57 0,0277 HIST2H2AA3; HIST2 histone cluster 2, H2aa3; histone cluster 2, H2aa4 -2,59 0,0054 ZNF594 zinc finger protein 594 -2,6 0,0409 FIGL1 figletin-like 1 -2,62 0,0101 MTMR2 myotubularin related protein 2 -2,62 0,0398 EFMH2 EF-hand domain (C-terminal) containing 2 -2,62 0,0265 OCLM Transcript identified by AceView, Entrez Gene ID(s) 10896 -2,63 6,69E-05 ZNF404 zinc finger protein 404 -2,63 0,0182 HRC2P3 hect domain and RLD 2 pseudogene 3 -2,63 0,0153 MINPP1 multiple inositol polyphosphate phosphatase 1 -2,66 0,0103 INTS1 Memczak2018 ANTISENSE, CDS, coding, INTERNAL best transcript NM_001080459 -2,66 0,0007 C1orf112 chromosome 1 open reading frame 112 -2,67 0,0053 SPATA25 spermatogenesis associated 25 -2,68 0,011 ZSWIM7 zinc finger, SWIM type containing 7 -2,7 0,0315 SLC26A2 solute carrier family 26 (anion exchanger), member 2 -2,71 0,0014 TMEM72 transmembrane protein 72 -2,73 0,0041 ZNF57 zinc finger protein 57 -2,73 0,0007 TP23A trans-golgi network vesicle protein 23 homolog A (S. cerevisiae) -2,75 0,0125 HIST3H2BD histone cluster 1, H2bd -2,76 0,0025 CRYGD crystallin gamma D -2,76 0,0195 FAM24B family with sequence similarity 24, member B -2,77 0,0064 AADAT aminoadipate aminotransferase -2,79 0,0162 CETN3 centrin 3 -2,81 0,028 ZNF883 zinc finger protein 883 -2,81 0,0038 CTB-75G16.3; MRM Transcript identified by AceView, Entrez Gene ID(s) 79922; novel transcript -2,81 0,0001 EGR1 early growth response 1 -2,86 0,0026 TTC33 tetratricopeptide repeat domain 33 -2,87 0,0064 QTRTD1 queuine tRNA-ribosyltransferase domain containing 1 -2,88 0,0042 RND1 Rho family GTPase 1 -2,89 0,0027 VP58 Transcript identified by AceView, Entrez Gene ID(s) 23355 -2,9 0,0176 TMEM139 transmembrane protein 139 -3 0,0298 GDDP3 glycerophosphodiester phosphodiesterase domain containing 3 -3,01 0,0122 TAF1A TATA box binding protein (TBP)-associated factor, RNA polymerase I, A, 48kDa -3,01 0,0004 CEP192 centrosomal protein 192kDa -3,09 0,0002 TSFD3 thio-sulfate sulfurtransferase (rhodanase)-like domain containing 3 -3,23 0,007 HIST3H2A histone cluster 3, H2a -3,35 0,0017 HIST3H3C histone cluster 1, H3c -3,35 0,0428 OCLN occludin -3,39 5,79E-05 HIST3H2BE Memczak2018 ANTISENSE, CDS, coding, upstream_start, UTR 3, UTR5 best transcript NM_003523 -3,94 0,0121 LHX4-AS1; ACBD6 LHX4 antisense RNA 1; acyl-CoA binding domain containing 6 -4,59 0,0016 NIPAL2 NIPA-like domain containing 2 -4,85 6,25E-05 NLGN1 neuroligin 1 -6,63 0,0018 OR2B2 olfactory receptor, family 2, subfamily B, member 2 -19,82 3,87E-05 HSP90B1 Transcript identified by AceView, Entrez Gene ID(s) 7184 </p>

Figure S12: Functional analysis of DEGs by Gene Ontology biological processes (GOBP) depicting 11 biological processes which are involved. Only upregulated but not downregulated genes were involved in these processes.

GO:BP		stats
Term name	Term ID	p _{adj}
regulation of response to stimulus	GO:0048583	6.932×10^{-6}
positive regulation of epidermal growth factor-activated rec...	GO:0045741	7.295×10^{-4}
ERBB2-EGFR signaling pathway	GO:0038134	1.270×10^{-2}
defense response to fungus	GO:0050832	1.951×10^{-2}
regulation of defense response	GO:0031347	2.751×10^{-2}
multicellular organismal process	GO:0032501	3.197×10^{-2}
cellular detoxification	GO:1990748	3.591×10^{-2}
regulation of metabolic process	GO:0019222	3.879×10^{-2}
homeostasis of number of cells	GO:0048872	4.119×10^{-2}
regulation of phosphate metabolic process	GO:0019220	4.988×10^{-2}
neutrophil aggregation	GO:0070488	4.995×10^{-2}