Pathway & Go term analysis

20190828

Zheng et al. 2017, T cell scRNAseqof Liver cancer. C4-CD8-LAYN cluster 20180905

Pathway analysis – Scheme (20180828)

Table S2 Zheng et al., 2017

List of signature genes in each cluster (C4-CD8-LAYN)

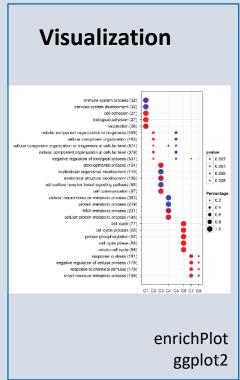
Extract EntrezID (274 genes)

	d(test)							
	5996 8486	58 5133	1493 100	663 201633				
> tes								
[1]	5996	84868	5133	1493	10663	201633		22822
[9]	4647	6348	953	939	678655	1846	143903	3122
[17]	4643	5552	9495	4929	84632	9760	1435	23406
[25]	1130	1757	2115	100287534	728558	3458	3805	55423
[33]	114614	9882	9452	10563	64759	116449	339789	79895
[41]	1390	64092	952	25861	3399	29851		84969
[49]	406947	5740	3108	5997	2597	7412	8302	64231
[57]	60489	4345	639	116841	3560	7453	3682	54
[65]	3398	7133	3902	9901	26191	8638	7462	84159
[73]	27333	160365	27115	7292	2530	2731		54602
[81]	55501	1509	285025	27350	4664	92906	3732	100528031
[89]	7128	5251	54438	5066	2280	677	3892	81
[97]	868	4046	356	1890	80727	100616438	25897	200316
[105]	3516	5763	3162	10018	924	7227	5873	64333
[113]	64218	79890	4689	164118	94240	7188	6310	10447
[121]	22876	6503	3428	894	10797	29988	8611	160851
[129]	79413	4739	5778	22806	970	967	2820	130589
[137]	5900	23231	27334	8832	140564	83982	79713	84959
[145]	7167	2537	8565	23062	10538	9260	9619	57162
[153]		100506190	10950	54806	9246	26031		9585
[161]	684	9746	3987	5781	2650	8536	4061	100130093
[169]	54625	57157	23710	4522	403	54434	7979	5583
[177]	11315	3984	65094	5359	100506755	10791	7037	10906
[185]	10421	1495	115361	5496	3430	7867	7633	6709
[193]	23741	80219	134429	836	23517	9766	8027	57414
[201]	5315	2762	6774	79718	9997	25824		2590
[209]	808	55320	81671	5573	4940	9144		50615
[217]	4599	308	9051	55500	10982	10459	2969	10725
[225]	3660	2182	10379		57835	29110		405
[233]	1499	399	219285	9158	5214	2625	2634	11243
[241]	5993	29959	54790	55601	11262	55072	100527963	4170
[249]	79415	57819	9446	9049	5686	3939	2130	10956
[257]	10095	847	3385	55608	9205	830	8775	9784
[265]	11164	6717	132160	1508	10241	282991	80344	4678
[273]	1460	58496						

Over-representation test (e.g. 250 genes – depending on geneset)

```
#...@organism
#...@ontology
                DisGeNET
#...@keytype
                 chr [1:274] "5996" "84868" "5133" "1493" "10663" "201633" "3604" "22822" "4647"
#...pvalues adjusted by 'BH' with cutoff <0.05
#...171 enriched terms found
 'data.frame': 171 obs. of 9 variables:
             : chr "umls:c0021400" "umls:c0024138" "umls:c0026896" "umls:c0409974" ...
 $ Description: chr "Influenza" "Lupus Erythematosus, Discoid" "Myasthenia Gravis" "Lupus Erythe
matosus" ...
 $ GeneRatio : chr
                    "28/250" "21/250" "16/250" "22/250"
 $ BgRatio : chr "494/17381" "316/17381" "181/17381" "356/17381" ...
             : num 6.47e-10 6.39e-09 8.06e-09 1.04e-08 2.08e-08 ...
 $ p.adjust : num 1.40e-06 5.62e-06 5.62e-06 5.62e-06 8.99e-06 ...
 $ qvalue : num 1.07e-06 4.31e-06 4.31e-06 4.31e-06 6.90e-06 ...
                    "1493/3604/6348/939/1757/3458/1390/2597/3682/8638/7292/25897/3162/2537/9246/
5781/4061/5359/3430/836/9997/4940/45"| _truncated_ "5133/1493/939/1435/3458/3805/10563/1390/298
51/639/7453/7133/26191/356/3428/970/84959/10950/6774/50615/2625" "5133/1493/3122/1435/3458/10563/
29851/406947/3560/26191/7128/5066/25897/9260/7979/836" "5133/1493/939/1435/3458/3805/10563/1390/9
52/29851/3108/639/7133/26191/356/3428/970/84959/10950/6774/50615/2625"...
             : int 28 21 16 22 20 21 24 13 24 10 ...
$ Count
#...Citation
 Guangchuang Yu, Li-Gen Wang, Yanyan Han and Qing-Yu He.
  clusterProfiler: an R package for comparing biological themes among
 gene clusters. OMICS: A Journal of Integrative Biology 2012, 16(5):284-287
                                                     DOSE – DisGeNET, DO
```

clusterProfiler - GO, KEGG



Limitation of the whole approach: ALL gene set analysis is knowledge-based. Novel genes and known genes with novel function will not be considered in the pathway analysis

Pathway analysis: over-representation analysis (ORA)

Over-representation analysis (ORA) - completed

- DisGenNET Disease
- DO ontology Disease
- KEGG Pathway
- GO ontology Cellular Component, Molecular Function, Biological Process

Limitations: ORA treats each gene equally! (No fold change or significance are considered for representation)

Analysis requires fold-changes or p-values as input (FCS)

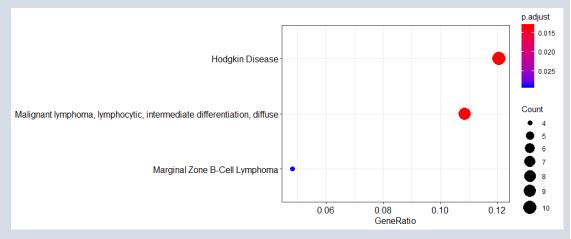
GSEA and others

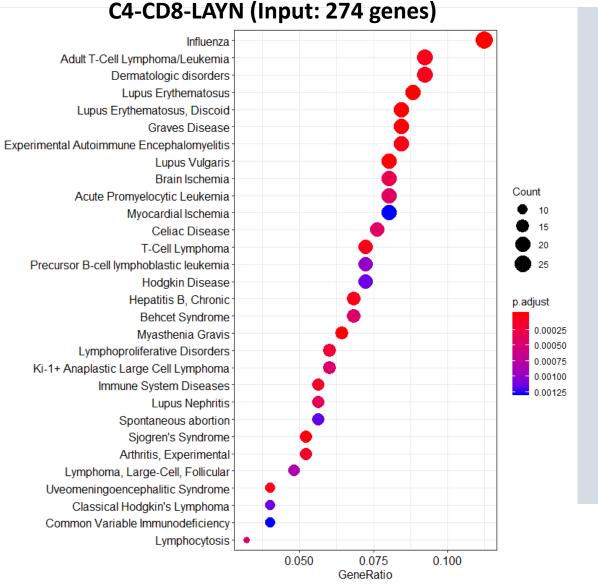
Comment: It does not require an arbitrary threshold for dividing expression data into significant and non-significant pool, but require whole gene expression dataset

Analysis: Over-representation test

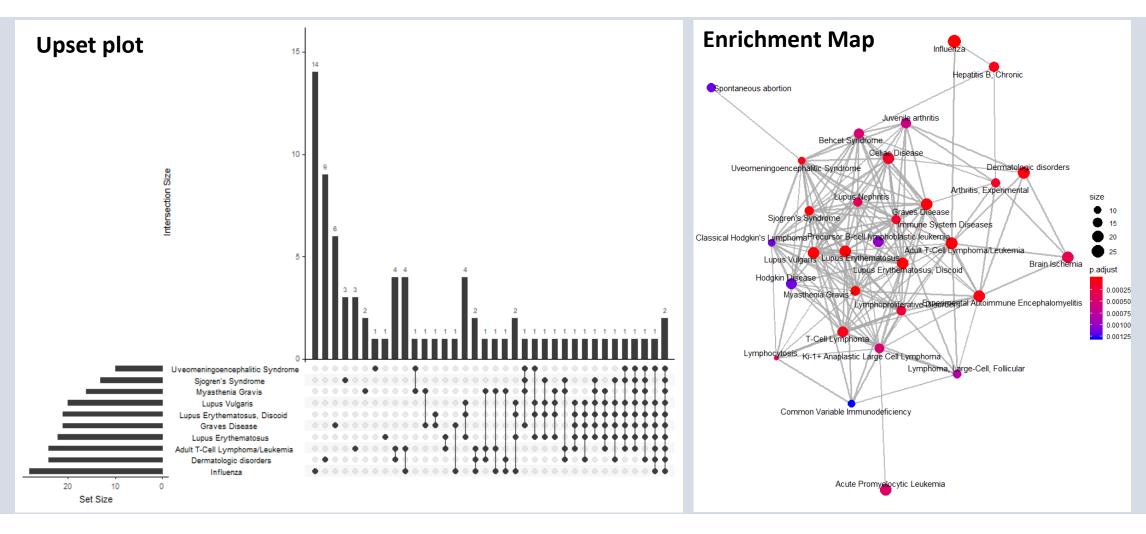


C1-CD8-LEF1 (Input: 101 genes)





Analysis: Visualization



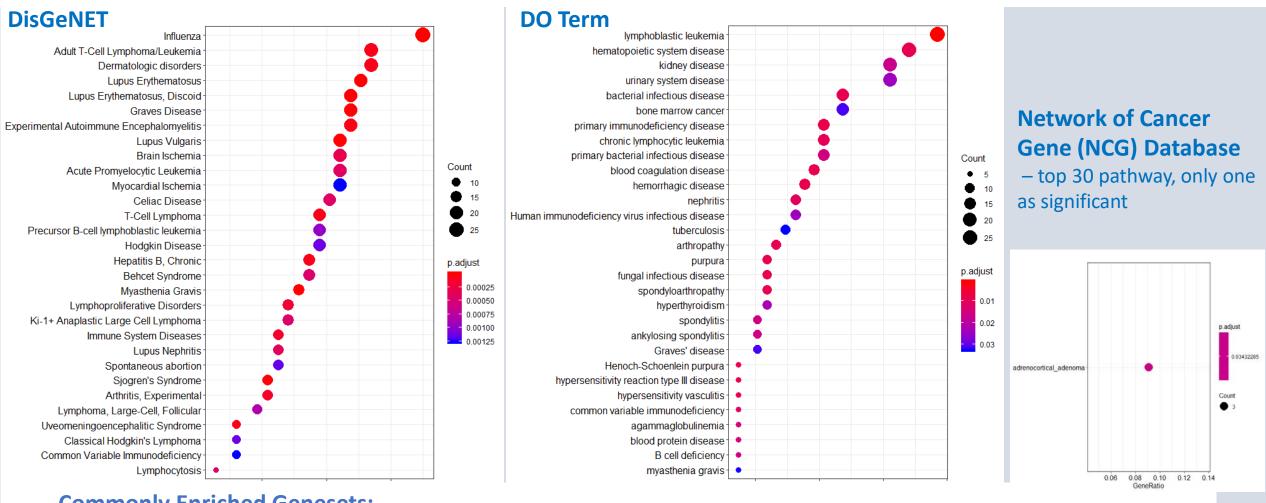
Analysis: Top Gene List (DisGeNet)

Description	GeneRatio pv	alue geneID
Influenza	28/250 6.	47E-10 CTLA4/TNFRSF9/CCL3/CD27/SARDH/IFNG/CREM/GAPDH/ITGAE/OASL/TNFSF4/RNF19A/HMOX1/IFI6/UBE2L6/PTPN11/LY6E/PLSCR1/IFI35/CASP3/SCO2/OAS3/MX1/IRF9/GBP1/GATA3/DDX60/CAT
Lupus Erythematosus, Discoid	21/250 6.	39E-09 PDCD1/CTLA4/CD27/CSF1/IFNG/KIR2DL4/CXCL13/CREM/ICOS/PRDM1/WARS/TNFRSF1B/PTPN22/FASLG/IFI16/CD70/UBASH3B/BTG3/STAT3/IL21R/GATA3
Myasthenia Gravis	16/250 8.	06E-09 PDCD1/CTLA4/HLA-DRA/CSF1/IFNG/CXCL13/ICOS/MIR155/IL2RB/PTPN22/TNFAIP3/PAM/RNF19A/PDLIM7/SEM1/CASP3
Lupus Erythematosus	22/250 1.	04E-08 PDCD1/CTLA4/CD27/CSF1/IFNG/KIR2DL4/CXCL13/CREM/CD38/ICOS/HLA-DMA/PRDM1/TNFRSF1B/PTPN22/FASLG/IFI16/CD70/UBASH3B/BTG3/STAT3/IL21R/GATA3
Lupus Vulgaris	20/250 2.	08E-08 PDCD1/CTLA4/CD27/CSF1/IFNG/KIR2DL4/CXCL13/CREM/ICOS/PRDM1/TNFRSF1B/PTPN22/FASLG/IFI16/CD70/UBASH3B/BTG3/STAT3/IL21R/GATA3
Graves Disease	21/250 4.	83E-08 PDCD1/CTLA4/CXCR6/IFNG/ITM2A/CXCL13/ICOS/WARS/ITGAE/TNFRSF1B/PTPN22/ARID5B/TNFAIP3/CBLB/FASLG/SEM1/STAT3/PRDX5/IL21R/RHOH/GATA3
Adult T-Cell Lymphoma/Leukemia	24/250 8.	09E-08 RGS1/PDCD1/CTLA4/CCL3/TOX/IFNG/CD38/ICOS/MIR155/VCAM1/IL2RB/TNFSF4/HMOX1/AHI1/CASP3/STAT3/OAS3/IL21R/IRF9/GATA3/MCL1/GSTO1/CAT/CSNK2B
Sjogren's Syndrome	13/250 8.	33E-08 PDCD1/CTLA4/IFNG/CXCL13/ID3/MIR155/PRDM1/PTPN22/TNFAIP3/FASLG/SPTAN1/STAT3/GTF2I
Dermatologic disorders	24/250 1.	20E-07 RGS1/CTLA4/CCL3/NR4A2/TOX/IFNG/MIR155/VCAM1/IL2RB/ID2/TNFRSF1B/PTPN22/FKBP1A/FASLG/RNF19A/HMOX1/IGFLR1/PELI1/PON2/MTHFD1/LIMK1/IRF2/GATA3/GSTO1
Uveomeningoencephalitic Syndrome	10/250 2.	16E-07 PDCD1/CTLA4/HLA-DRA/IFNG/KIR2DL4/PTPN22/TNFAIP3/TRAF5/STAT3/GATA3
Experimental Autoimmune Encephalomyelitis	21/250 2.	25E-07 HAVCR2/CTLA4/TNFRSF9/CCL3/ENTPD1/NR4A2/IFNG/CXCL13/MIR155/VCAM1/IL2RB/TNFRSF1B/CBLB/RNF19A/SEMA4A/CASP3/STAT3/PRDX5/GATA3/GSTO1/CAT
Hepatitis B, Chronic	17/250 3.	41E-07 HAVCR2/PDCD1/CTLA4/TNFRSF9/IFNG/KIR2DL4/KLRC4/APOBEC3G/WARS/ACP5/OASL/FASLG/OAS3/MX1/IRF2/IRF9/CAT
T-Cell Lymphoma	18/250 3.	68E-07 PDCD1/CTLA4/CD27/IFNG/CXCL13/ID3/MIR155/PRDM1/IL2RB/ID2/BCL2L11/CCND2/CD70/PDLIM7/STAT3/CTNNB1/GATA3/TET2
Immune System Diseases	14/250 8.	60E-07 HAVCR2/PDCD1/CTLA4/ENTPD1/IFNG/ICOS/PTPN22/TNFAIP3/RAB27A/IKZF3/BATF/STAT3/GATA3/TET2
Arthritis, Experimental	13/250 9.	07E-07 HAVCR2/CTLA4/CCL3/NR4A2/IFNG/TNFRSF1B/PAM/CBLB/FASLG/STAT3/ACSL4/CAT/CTSB
Celiac Disease	20/250 9.	88E-07 RGS1/PDCD1/CTLA4/TNFRSF9/HLA-DRA/IFNG/KIR2DL4/CD38/ICOS/IL2RB/PTPN22/TNFSF4/TNFAIP3/ZFP36L1/FASLG/PON2/PARK7/TFRC/GATA3/CTSB
Lymphoproliferative Disorders	15/250 1.	56E-06 CTLA4/CD27/IFNG/CXCL13/CD38/MIR155/PRDM1/IL2RB/TNFRSF1B/TNFAIP3/CBLB/FASLG/BCL2L11/PDLIM7/STAT3
Brain Ischemia	20/250 2.	56E-06 CTLA4/CCL3/TOX/PTGIS/VCAM1/ITGAE/TNFRSF1B/FASLG/HMOX1/PON2/MTHFD1/PRKCH/LIMK1/CASP3/STAT3/GALNT2/GSTO1/CAT/CTSB/LY6G5B
Lupus Nephritis	14/250 3.	25E-06 PDCD1/CSF1/IFNG/CXCL13/VCAM1/TNFSF4/TNFAIP3/FASLG/UBASH3B/LY6E/CASP3/STAT3/MX1/GATA3
Acute Promyelocytic Leukemia	20/250 4.	08E-06 SRGN/CSF1/IFNG/CD38/ASB2/ID2/FASLG/ATXN1/UBE2L6/PLSCR1/TFRC/CASP3/STAT3/TBL1XR1/PRDX5/PRKAR1A/IRF9/TET2/SP140/MCL1
Lymphocytosis	8/250 4.	58E-06 CTLA4/CD27/IFNG/CD38/MIR155/ITGAE/FASLG/PDLIM7
Behcet Syndrome	17/250 5.	05E-06 PDCD1/CTLA4/IFNG/KIR2DL4/KLRC4/TNFRSF1B/PTPN22/TNFAIP3/HMOX1/TRAF5/IFI16/UBASH3B/PHTF2/STAT3/PSTPIP1/GATA3/CAT
Ki-1+ Ananlactic Large Cell Lumphoma	15/250 5	10E_06 IENG/MIR155 /DRDM1 / ID2 /TNERSE1R /EASI G/RCI 2L11 /DDI IM2 /DTDM11 /CASD2/STAT2 /CALM2 / IL 21R /GATA2 /MCL1

Caution 1: Only "upregulated genes" were defined as signature genes to start with, but loss of function mutation etc could define select gene as "geneset"

Caution 2 : Each pathway was treated independently – a gene can function in multiple pathway so identified pathways were frequently overlapped → Obscure true source of signals

Analysis: DO term analysis



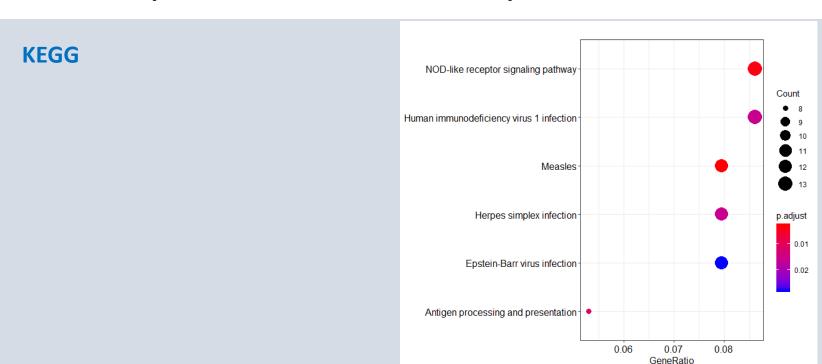
Commonly Enriched Genesets:

Infectious disease, Leukemia/Lymphoma, Autoimmune disease, Graves Disease/hyperthyroidism, Blood Vessle (Ischemia ←→Purpura/hemorrhagic/Behcet Syndrome)

Analysis: Top Gene List (DO term)

ID	Description	GeneRatio	p.adjust geneID
DOID:1037	lymphoblastic leukemia	26/169	0.001016 CCL3/IFNG/CXCL13/CD38/IL2RB/LAT2/ARID5B/CLECL1/PDE7B/TNFAIP3/ZFP36L1/FASLG/BCL2L11/CCND2/IKZF3/CD70/PTPN11/TFRC/PKM/TBL1XR1/IL21R/ARNT/RHOH/MCL1/LDHA/CSNK2B
DOID:612	primary immunodeficiency disease	14/169	0.008202 PDCD1/CTLA4/IFNG/ICOS/APOBEC3G/TNFRSF1B/FASLG/APOBEC3F/NCF4/CD70/CD63/BST2/PRKAR1A/NFAT5
DOID:11123	Henoch-Schoenlein purpura	5/169	0.008202 CTLA4/PTPN22/TNFSF4/HMOX1/CAT
DOID:1557	hypersensitivity reaction type III disease	5 5/169	0.008202 CTLA4/PTPN22/TNFSF4/HMOX1/CAT
DOID:9809	hypersensitivity vasculitis	5/169	0.008202 CTLA4/PTPN22/TNFSF4/HMOX1/CAT
DOID:2213	hemorrhagic disease	12/169	0.008202 CTLA4/HLA-DRA/IFNG/PTPN22/TNFSF4/FASLG/HMOX1/CASP3/CTNNB1/TET2/LDHA/CAT
DOID:1247	blood coagulation disease	13/169	0.008473 CTLA4/HLA-DRA/IFNG/PTPN22/TNFSF4/FASLG/HMOX1/PTPN11/CASP3/CTNNB1/TET2/LDHA/CAT
DOID:3326	purpura	8/169	0.008473 CTLA4/IFNG/PTPN22/TNFSF4/FASLG/HMOX1/CASP3/CAT
DOID:74	hematopoietic system disease	23/169	0.009264 CTLA4/HLA-DRA/IFNG/CD38/ICOS/VCAM1/TNFRSF1B/PTPN22/TNFSF4/FASLG/HMOX1/CD70/PTPN11/TFRC/SPTAN1/CASP3/PRKAR1A/NFAT5/CTNNB1/GATA3/TET2/LDHA/CAT
DOID:104	bacterial infectious disease	16/169	0.009264 PDCD1/CTLA4/CXCR6/CCL3/CD27/IFNG/CD38/ICOS/PRDM1/TNFRSF1B/PTPN22/CBLB/FASLG/BCL2L11/STAT3/MCL1
DOID:1564	fungal infectious disease	8/169	0.009264 PDCD1/CTLA4/CXCR6/IFNG/TNFRSF1B/PTPN22/STAT3/ICAM3
DOID:381	arthropathy	9/169	0.009264 RGS1/PDCD1/CTLA4/CSF1/IFNG/ICOS/PTGIS/TNFRSF1B/STAT3
DOID:1123	spondyloarthropathy	8/169	0.009264 RGS1/PDCD1/CTLA4/IFNG/ICOS/PTGIS/TNFRSF1B/STAT3
DOID:1040	chronic lymphocytic leukemia	14/169	0.00968 CCL3/IFNG/CXCL13/CD38/PDE7B/TNFAIP3/ZFP36L1/FASLG/BCL2L11/TFRC/IL21R/RHOH/MCL1/CSNK2B
DOID:10952	nephritis	11/169	0.010447 PDCD1/MYO1E/CSF1/IFNG/CXCL13/VCAM1/TNFSF4/ACTN4/FASLG/CASP3/GATA3
DOID:12177	common variable immunodeficiency	5/169	0.011504 CTLA4/ICOS/TNFRSF1B/CD70/PRKAR1A
DOID:2583	agammaglobulinemia	5/169	0.01414 CTLA4/ICOS/TNFRSF1B/CD70/PRKAR1A
DOID:620	blood protein disease	5/169	0.01414 CTLA4/ICOS/TNFRSF1B/CD70/PRKAR1A
DOID:0050338	8 primary bacterial infectious disease	14/169	0.015208 PDCD1/CTLA4/CCL3/CD27/IFNG/CD38/ICOS/PRDM1/TNFRSF1B/PTPN22/CBLB/FASLG/BCL2L11/MCL1
DOID:6590	spondylitis	7/169	0.015915 RGS1/PDCD1/CTLA4/IFNG/PTGIS/TNFRSF1B/STAT3
DOID:2115	B cell deficiency	5/169	0.015915 CTLA4/ICOS/TNFRSF1B/CD70/PRKAR1A
DOID:7147	ankylosing spondylitis	7/169	0.015915 RGS1/PDCD1/CTLA4/IFNG/PTGIS/TNFRSF1B/STAT3
DOID:557	kidney disease	21/169	0.017219 PDCD1/MYO1E/CSF1/IFNG/CXCL13/RGS2/VCAM1/ACP5/TNFRSF1B/TNFSF4/ACTN4/FASLG/RBPJ/HMOX1/AHI1/LIMS1/TFRC/CASP3/STAT3/GATA3/ICAM3
DOID:7998	hyperthyroidism	8/169	0.022529 CTLA4/CXCR6/TNFRSF9/IFNG/TNFRSF1B/PTPN22/FASLG/PRDX5
DOID:526	Human immunodeficiency virus infection	c 11/169	0.024254 PDCD1/CXCR6/CCL3/CD27/CSF1/IFNG/CXCL13/VCAM1/APOBEC3G/IL2RB/FASLG
DOID:18	urinary system disease	21/169	0.024254 PDCD1/MYO1E/CSF1/IFNG/CXCL13/RGS2/VCAM1/ACP5/TNFRSF1B/TNFSF4/ACTN4/FASLG/RBPJ/HMOX1/AHI1/LIMS1/TFRC/CASP3/STAT3/GATA3/ICAM3
DOID:12361	Graves' disease	7/169	0.031455 CTLA4/CXCR6/TNFRSF1B/PTPN22/FASLG/PRDX5
DOID:4960	bone marrow cancer	16/169	0.031469 CCL3/SRGN/IFNG/VCAM1/PRDM1/ACP5/LAG3/FASLG/BCL2L11/CCND2/PTPN11/STAT3/CTNNB1/TET2/MCL1/LDHA
DOID:437	myasthenia gravis	5/169	0.032656 PDCD1/CTLA4/HLA-DRA/CXCL13/PTPN22

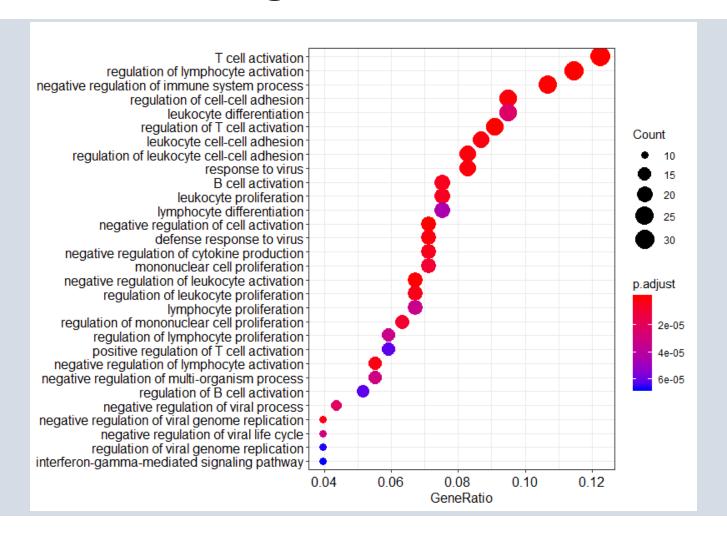
Analysis: KEGG analysis



ID	Description	GeneRatio	o p.adjust	qvalue	geneID	Count
hsa05162	Measles	12/151	0.003005	0.002636	6 IFNG/IL2RB/TNFAIP3/CBLB/FASLG/CCND2/STAT3/OAS3/MX1/IRF9/TBK1/CSNK2B	12
hsa04621	NOD-like receptor signaling pathway	13/151	0.003707	0.003251	1 TNFAIP3/TRAF5/IFI16/GABARAPL1/GBP4/OAS3/PSTPIP1/IRF9/GBP1/TBK1/GBP2/RNF31/CTSB	13
hsa04612	Antigen processing and presentation	8/151	0.011375	0.009978	8 HLA-DRA/IFNG/KIR2DL4/HLA-DMA/KLRC4/PSME2/RFX5/CTSB	8
hsa05170	Human immunodeficiency virus 1 infection	13/151	0.015951	0.013997	2 APOBEC3G/TNFRSF1B/APOBEC3C/FASLG/APOBEC3F/TRAF5/APOBEC3D/BST2/LIMK1/CASP3/CALM3/TBK1/MAP2K3	13
hsa05168	Herpes simplex infection	12/151	0.015951	0.013997	2 HLA-DRA/IFNG/HLA-DMA/FASLG/TRAF5/PTPN11/CASP3/OAS3/GTF2I/IRF9/TBK1/CSNK2B	12
hsa05169	Epstein-Barr virus infection	12/151	0.028047	0.024603	3 ENTPD1/HLA-DRA/IFNG/CD38/TNFAIP3/RBPJ/TRAF5/SEM1/STAT3/TBK1/MAP2K3/CSNK2B	12

C4-CD8-LAYN (Input: 274 genes)

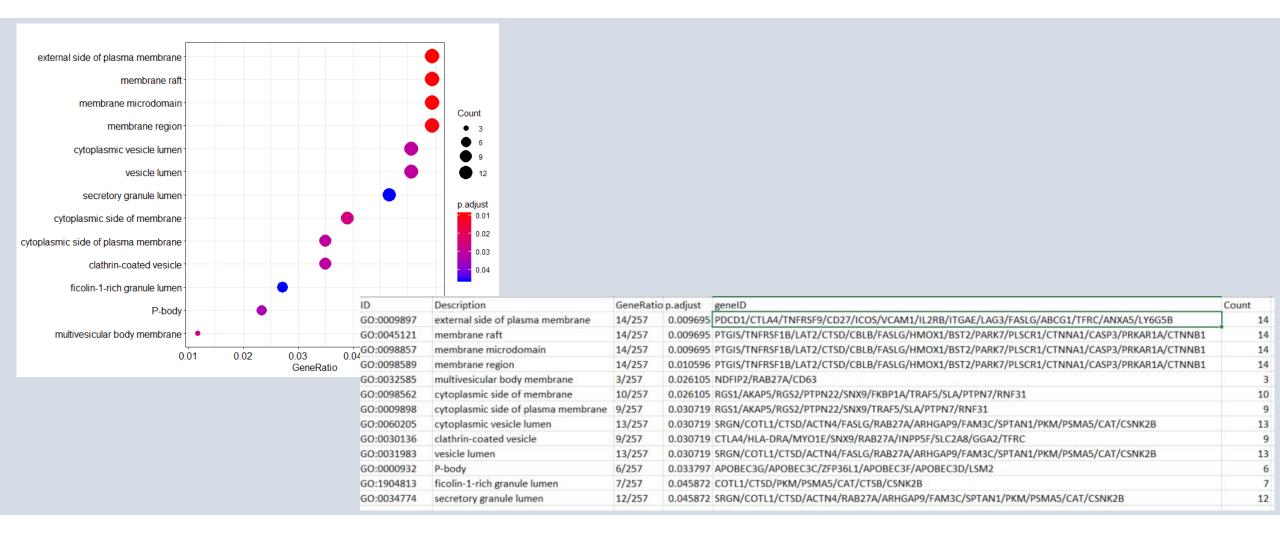
Analysis: GO – Biological Process



Analysis: GO – Biological Process

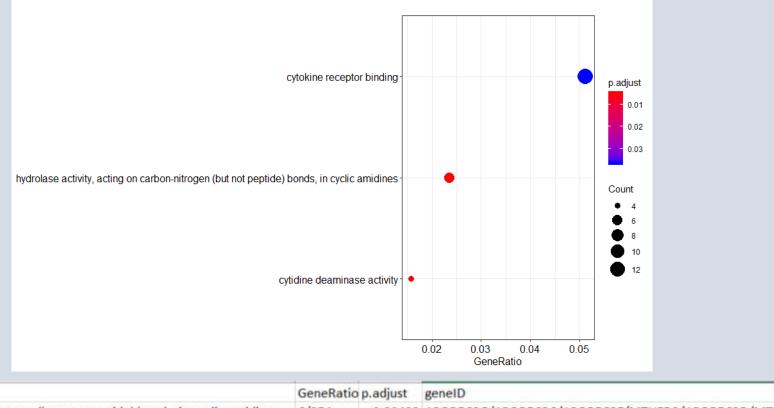
ID	Description	GeneRatio	p.adjust geneID
GO:0042110	T cell activation	31/253	3.49E-09 HAVCR2/PDCD1/CTLA4/TIGIT/CD27/HLA-DRA/IFNG/SIRPG/ICOS/MIR155/VCAM1/PRDM1/LAG3/PTPN22/CLECL1/TNFSF4/FKBP1A/ZFP36L1/CD7/RAB27A/SEMA4A/BATF/PELI1/PTPN11/TFRC/CASP3/STAT3/PRKAR1A/CTNNB1/RHOH/GATA3
GO:0051249	regulation of lymphocyte activation	29/253	3.69E-08 HAVCR2/PDCD1/CTLA4/TIGIT/CD27/HLA-DRA/IFNG/SIRPG/SAMSN1/CD38/ICOS/MIR155/VCAM1/PRDM1/ID2/LAG3/PTPN22/CLECL1/TNFSF4/TNFAIP3/ZFP36L1/IKZF3/PELI1/PTPN11/TFRC/CASP3/PRKAR1A/CTNNB1/GATA3
GO:0050866	negative regulation of cell activation	18/253	7.36E-08 HAVCR2/CTLA4/TIGIT/SAMSN1/MIR155/CD200/PRDM1/ID2/LAG3/PTPN22/TNFSF4/TNFAIP3/HMOX1/CD84/UBASH3B/PELI1/CASP3/PRKAR1A
GO:0002695	negative regulation of leukocyte activation	17/253	7.45E-08 HAVCR2/CTLA4/TIGIT/SAMSN1/MIR155/CD200/PRDM1/ID2/LAG3/PTPN22/TNFSF4/TNFAIP3/HMOX1/CD84/PELI1/CASP3/PRKAR1A
GO:0002683	negative regulation of immune system process	27/253	7.87E-08 HAVCR2/PDCD1/CTLA4/TIGIT/CCL3/SAMSN1/MIR155/CD200/PRDM1/ID2/LAG3/PTPN22/TNFSF4/TNFAIP3/ZFP36L1/HMOX1/IFI16/CD84/UBASH3B/PELI1/BST2/PARP14/TRAFD1/CASP3/PRKAR1A/GBP1/CTNNB1
GO:0050863	regulation of T cell activation	23/253	1.57E-07 HAVCR2/PDCD1/CTLA4/TIGIT/CD27/HLA-DRA/IFNG/SIRPG/ICOS/MIR155/VCAM1/PRDM1/LAG3/PTPN22/CLECL1/TNFSF4/PELI1/PTPN11/TFRC/CASP3/PRKAR1A/CTNNB1/GATA3
GO:0022407	regulation of cell-cell adhesion	24/253	1.06E-06 HAVCR2/PDCD1/CTLA4/TIGIT/CD27/HLA-DRA/IFNG/SIRPG/CXCL13/ICOS/MIR155/VCAM1/LAG3/PTPN22/CLECL1/TNFSF4/UBASH3B/PELI1/PTPN11/TFRC/CASP3/PRKAR1A/MAD2L2/GATA3
GO:1903037	regulation of leukocyte cell-cell adhesion	21/253	1.26E-06 HAVCR2/PDCD1/CTLA4/TIGIT/CD27/HLA-DRA/IFNG/SIRPG/ICOS/MIR155/VCAM1/LAG3/PTPN22/CLECL1/TNFSF4/PELI1/PTPN11/TFRC/CASP3/PRKAR1A/GATA3
GO:0007159	leukocyte cell-cell adhesion	22/253	1.33E-06 HAVCR2/PDCD1/CTLA4/TIGIT/CD27/HLA-DRA/IFNG/SIRPG/ICOS/MIR155/VCAM1/LAG3/PTPN22/CLECL1/TNFSF4/PELI1/PTPN11/GCNT1/TFRC/CASP3/PRKAR1A/GATA3
GO:0051607	defense response to virus	18/253	1.46E-06 LYST/IFNG/APOBEC3G/OASL/APOBEC3C/TNFAIP3/APOBEC3F/IF116/APOBEC3D/BST2/PLSCR1/OAS3/MX1/IRF2/IRF9/GBP1/TBK1/DDX60
GO:0009615	response to virus	21/253	1.46E-06_LYST/IFNG/APOBEC3G/OASL/TNFSF4/APOBEC3C/TNFAIP3/APOBEC3F/BCL2L11/IF116/APOBEC3D/BST2/PLSCR1/OAS3/MX1/IRF2/IRF9/GBP1/TBK1/GATA3/DDX60
GO:0045071	negative regulation of viral genome replication	10/253	1.58E-06 APOBEC3G/OASL/APOBEC3F/IFI16/APOBEC3D/BST2/PLSCR1/OAS3/MX1
GO:0051250	negative regulation of lymphocyte activation	14/253	1.88E-06 HAVCR2/CTLA4/TIGIT/SAMSN1/MIR155/PRDM1/ID2/LAG3/PTPN22/TNFSF4/TNFAIP3/PELI1/CASP3/PRKAR1A
GO:0070663	regulation of leukocyte proliferation	17/253	3.01E-06 HAVCR2/CTLA4/CSF1/CD38/MIR155/VCAM1/PRDM1/PTPN22/CLECL1/TNFSF4/TNFAIP3/IKZF3/PELI1/TFRC/CASP3/PRKAR1A/CTNNB1
GO:0042113	B cell activation	19/253	3.81E-06 CTLA4/CD27/ITM2A/SAMSN1/CD38/VCAM1/PRDM1/ID2/LAT2/TNFSF4/TNFAIP3/ZFP36L1/RBPJ/IKZF3/BATF/PELI1/BST2/TFRC/CASP3
GO:0001818	negative regulation of cytokine production	18/253	3.81E-06 HAVCR2/TIGIT/TNFRSF9/SRGN/IFNG/MIR155/ACP5/LAG3/PTPN22/TNFSF4/TNFAIP3/HMOX1/CD84/UBE2L6/BST2/GBP1/TBK1/GATA3
GO:0070661	leukocyte proliferation	19/253	4.59E-06 HAVCR2/CTLA4/CSF1/CD38/MIR155/VCAM1/PRDM1/PTPN22/CLECL1/TNFSF4/TNFAIP3/FKBP1A/IKZF3/PELI1/TFRC/CASP3/PRKAR1A/TBK1/CTNNB1
GO:0032944	regulation of mononuclear cell proliferation	16/253	7.60E-06 HAVCR2/CTLA4/CSF1/CD38/MIR155/VCAM1/PRDM1/PTPN22/CLECL1/TNFSF4/IKZF3/PELI1/TFRC/CASP3/PRKAR1A/CTNNB1
GO:0032943	mononuclear cell proliferation	18/253	9.03E-06 HAVCR2/CTLA4/CSF1/CD38/MIR155/VCAM1/PRDM1/PTPN22/CLECL1/TNFSF4/FKBP1A/IKZF3/PELI1/TFRC/CASP3/PRKAR1A/TBK1/CTNNB1
GO:0048525	negative regulation of viral process	11/253	2.12E-05 CCL3/APOBEC3G/OASL/APOBEC3F/IFI16/APOBEC3D/BST2/PLSCR1/OAS3/MX1
GO:0002521	leukocyte differentiation	24/253	2.33E-05 CTLA4/CCL3/CD27/CSF1/IFNG/ITM2A/MIR155/VCAM1/PRDM1/ID2/PTPN22/TNFSF4/ZFP36L1/RBPJ/SEMA4A/IFI16/IKZF3/UBASH3B/BATF/TFRC/STAT3/CTNNB1/RHOH/GATA3
GO:1903901	negative regulation of viral life cycle	10/253	3.08E-05 APOBEC3G/OASL/APOBEC3F/IFI16/APOBEC3D/BST2/PLSCR1/OAS3/MX1
GO:0043901	negative regulation of multi-organism process	14/253	3.08E-05 HAVCR2/CCL3/APOBEC3G/PRDM1/OASL/APOBEC3C/TNFAIP3/APOBEC3D/BST2/PLSCR1/OAS3/MX1
GO:0050670	regulation of lymphocyte proliferation	15/253	3.33E-05 HAVCR2/CTLA4/CD38/MIR155/VCAM1/PRDM1/PTPN22/CLECL1/TNFSF4/IKZF3/PELI1/TFRC/CASP3/PRKAR1A/CTNNB1
GO:0046651	lymphocyte proliferation	17/253	3.33E-05 HAVCR2/CTLA4/CD38/MIR155/VCAM1/PRDM1/PTPN22/CLECL1/TNFSF4/FKBP1A/IKZF3/PELI1/TFRC/CASP3/PRKAR1A/TBK1/CTNNB1
GO:0030098	lymphocyte differentiation	19/253	4.45E-05 CTLA4/CD27/IFNG/ITM2A/MIR155/VCAM1/PRDM1/ID2/PTPN22/TNFSF4/ZFP36L1/RBPJ/SEMA4A/IKZF3/BATF/STAT3/CTNNB1/RHOH/GATA3
GO:0050870	positive regulation of T cell activation	15/253	6.17E-05 HAVCR2/PDCD1/CTLA4/CD27/HLA-DRA/IFNG/SIRPG/ICOS/MIR155/VCAM1/CLECL1/TNFSF4/PTPN11/TFRC/GATA3
GO:0050864	regulation of B cell activation	13/253	6.17E-05 CTLA4/CD27/SAMSN1/CD38/PRDM1/ID2/TNFSF4/TNFAIP3/ZFP36L1/IKZF3/PELI1/TFRC/CASP3
GO:0045069	regulation of viral genome replication	10/253	6.66E-05 APOBEC3G/OASL/APOBEC3F/IFI16/APOBEC3D/BST2/PLSCR1/OAS3/MX1
GO:0060333	interferon-gamma-mediated signaling pathway	10/253	6.71E-05 HLA-DRA/IFNG/VCAM1/OASL/PARP14/OAS3/IRF2/IRF9/GBP1/GBP2
GO:0060337	type I interferon signaling pathway	10/253	6.71E-05 OASL/IFI6/BST2/PTPN11/IFI35/OAS3/MX1/IRF2/IRF9/GBP2
GO:0071357	cellular response to type I interferon	10/253	6.71E-05 OASL/IFI6/BST2/PTPN11/IFI35/OAS3/MX1/IRF2/IRF9/GBP2
GO:0034341	response to interferon-gamma	14/253	7.74E-05 CCL3/HLA-DRA/IFNG/GAPDH/VCAM1/OASL/BST2/PARP14/OAS3/IRF2/IRF9/GBP1/GBP2/CALCOCO2
GO:0051251	positive regulation of lymphocyte activation	18/253	7.84E-05 HAVCR2/PDCD1/CTLA4/CD27/HLA-DRA/IFNG/SIRPG/CD38/ICOS/MIR155/VCAM1/PRDM1/CLECL1/TNFSF4/PELI1/PTPN11/TFRC/GATA3
GO:0019079	viral genome replication	11/253	8.59E-05 CXCR6/APOBEC3G/OASL/APOBEC3F/IFI16/APOBEC3D/BST2/PLSCR1/OAS3/MX1

Analysis: GO – Cellular component



C4-CD8-LAYN (Input: 274 genes)

Analysis: GO – molecular function



ID	Description	GeneRatio	p.adjust	geneID
GO:0016814	hydrolase activity, acting on carbon-nitrogen (but not peptide) bonds, in cyclic amidines	6/254	0.00492	APOBEC3G/APOBEC3C/APOBEC3F/MTHFD2/APOBEC3D/MTHFD1
GO:0004126	cytidine deaminase activity	4/254	0.00492	APOBEC3G/APOBEC3F/APOBEC3D
GO:0005126	cytokine receptor binding	13/254	0.036655	CCL3/CSF1/IFNG/CXCL13/TNFSF4/FKBP1A/FASLG/TRAF5/CD70/YARS/CASP3/STAT3/GATA3

C4-CD8-LAYN (Input: 274 genes)

Ongoing pathway Analysis (Future plan)

- Advanced analysis with fold changes & p-values inclusion
- Modified GSEA analysis using partial list of gene set might be doable
 - Without triplicate input? Maybe I can manually add calculated statistics.
 - I have only partial list of gene expression How rank is determined?
- Sorting genes expressed on cell surfaces
- Came up with potential target to follow