

Supplementary Materials

An activated CD8⁺ T cell-based signature stratifies prognosis and suggests differential sensitivity to immune checkpoint inhibitors in hepatocellular carcinoma

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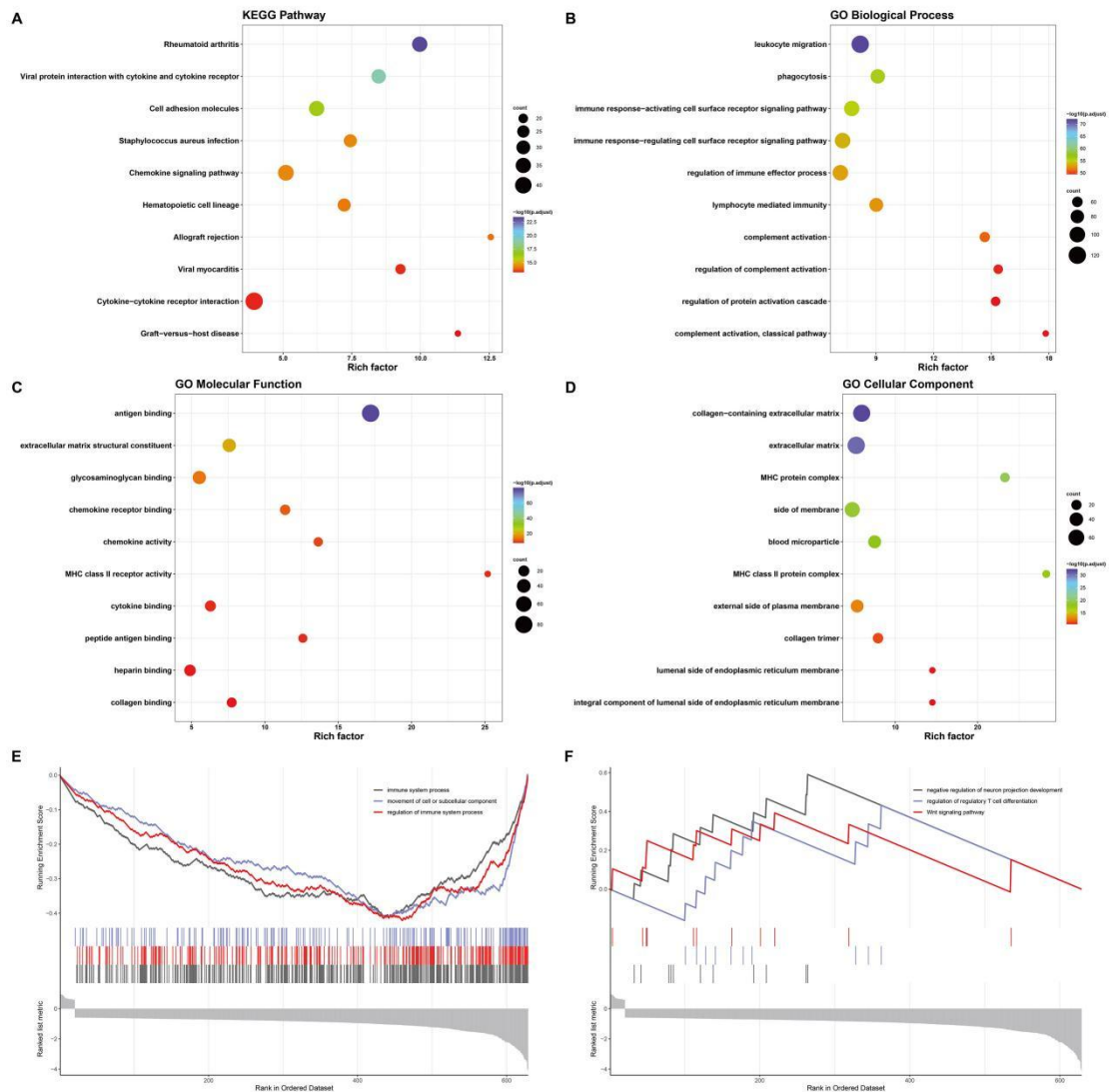
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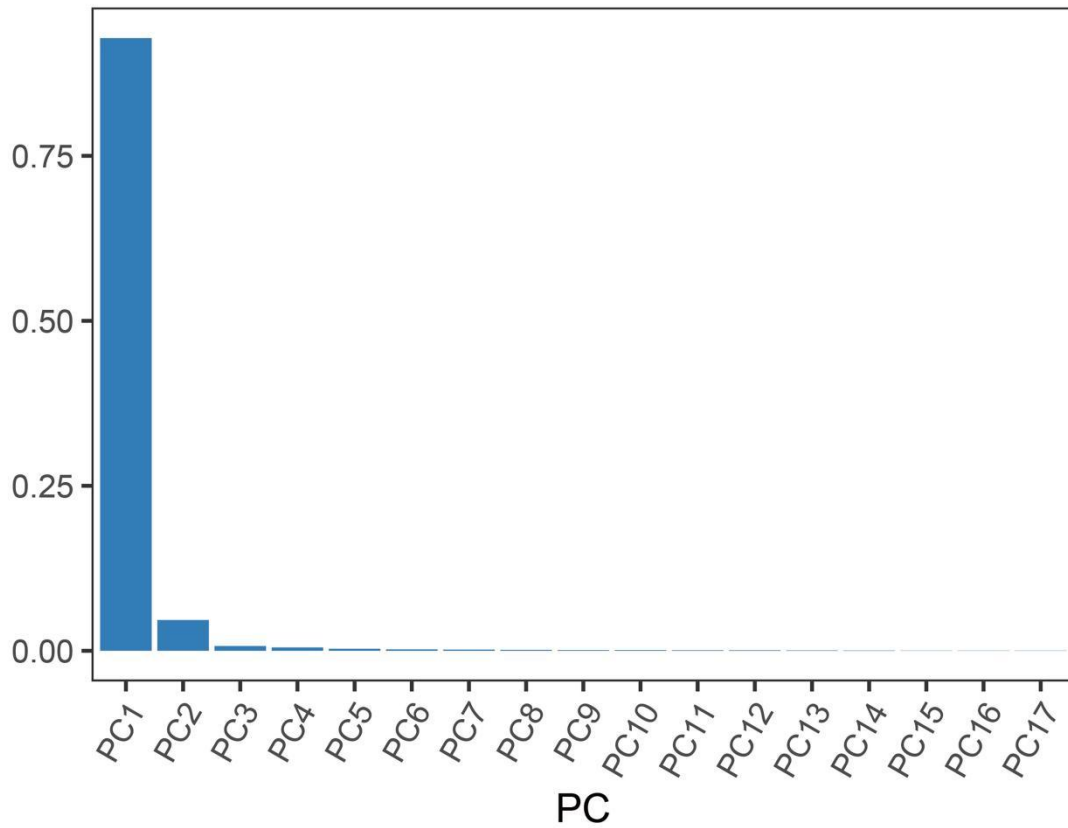
[#]These authors contributed equally to this work.

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(C) Significantly enriched GO terms in molecular function; (D) Significantly enriched GO terms in cellular component. KEGG: Kyoto Encyclopedia of Genes and Genomes; GO: gene ontology.



Supplementary Figure 3. Pathway enrichment analysis of 648 differentially expressed genes between clusters. (A) KEGG pathways enriched among the 648 DEGs; (B) Significantly enriched GO terms in biological process; (C) Significantly enriched GO terms in molecular function; (D) Significantly enriched GO terms in cellular component; (E) GSEA plots in Cluster 1; (F) GSEA plots in Cluster 2. KEGG: Kyoto Encyclopedia of Genes and Genomes; DEGs: differentially expressed genes; GO: gene ontology; GSEA: gene set enrichment analysis.



Supplementary Figure 4. Variance explained by each principal component in PCA.
PCA: Principal component analysis.

Supplementary Table 1. Clinical characteristics of HCC patients in the analyzed datasets

Parameter	Subtype	TCGA	GSE76427	GSE14520
Gender	female	118	22	30
	male	245	93	190
	na	-	-	-
Age	> 60	190	67	39
	< = 60	173	48	181
	na	-	-	-
Stage	i	170	55	93
	ii	84	35	77

	iii	81	21	48
	iv	4	3	-
	na	24	1	2
M	M0	262	-	-
	M1+MX	101	-	-
N	N0	246	-	-
	N1+NX	116	-	-
	na	1	-	-
T	T1	180	-	-
	T2	91	-	-
	T3	76	-	-
	T4	13	-	-
	TX	1	-	-
	na	2	-	-
Race	American indian or Alaska native	1	-	-
	Asian	155	-	-
	Black or african American	17	-	-
	White	180	-	-
	na	10	-	-

HCC: Hepatocellular carcinoma; TCGA: the Cancer Genome Atlas.

Supplementary Table 2. List of genes assigned to each WGCNA module

Module	Gene
Blue	NFYA
	SEMA3F
	CFLAR
	SARM1
	DHX33
	MAP3K14

IFRD1
ADIPOR2
IL32
RANBP9
HFE
ZMYND11
RABEP1
PIAS1
GOPC
KITLG
MAPK9
BCAR1
PIK3CB
EIF2AK2
ATG5
CASP8
GNAI3
MAP2K4
ELAVL1
NFYC
NEO1
RHOA
FGFR3
OTUD5
NEDD4
CDC42
TRIB2
TFRC
NFATC3

SMARCE1
EED
PLXNA2
MAP2K7
PIAS2
PIK3C3
ITCH
TP73
XRCC5
MKNK1
DDX1
SMARCA2
MID2
CNOT4
HSP90AA1
GSK3B
CYLD
REST
TXLNA
HACE1
MAP3K4
ATG16L1
DNM1L
KHSRP
MAVS
P2RX7
CMTM1
OTUB2
RCOR1

MUL1
CMTM6
RNF31
SNAP23
TGFB2
LRRFIP2
MAP3K1
PSMD5
HSP90AB1
SIRT1
JAK2
ABL1
NRP1
CRKL
MAPK1
DDX17
MYH9
SOS2
PSMC6
HIF1A
DICER1
PSMC1
RPS6KA5
YY1
TRPC4AP
NFATC2
PSMD10
MID1
XIAP

NFAT5
CTCF
TCEB2
UBE2W
IKBKB
TNFRSF10A
HNRNPL
AKT2
PIAS4
TYK2
CDK6
ZC3HAV1
LMBR1
TAX1BP1
RARRES2
AHR
CORO2A
TGFB1
MAPK8
PPP3CB
BMP1A
MAP3K8
PSMD3
TRIM37
AKAP10
PSMD11
EFTUD2
MAP2K6
VTN

NFKB1
GAB1
BIRC2
CBL
CAMKK2
DUSP16
ULBP1
PPARD
MAPK14
ASCC3
VEGFA
ERBB2IP
SEMA5A
BRD8
PRLR
NR3C1
TRIM23
KPNA1
KAT2B
USP4
FXR1
CBLB
PLXNA1
ACVR2B
PSMD14
CCDC88A
LANCL1
IL1R1
SOS1

ATF2
NFE2L2
CACYBP
TRIM62
MFN2
NR5A2
LGALS8
CD46
MPL
DR1
TNFSF4
NENF
APOA1
CREB1
TNFAIP3
FOXO3
CTNNAL1
TRIM32
ELF1
NR2C1
TNFRSF10B
TRIM25
PIK3CA
ACVR2A
PTGFR
TRIM24
ECD
C4BPB
PLCG1

NUP153
LRRFIP1
AHNAK
ABCC4
BMP4
THRA
SBDS
BECN1
NUP214
CANX
FGFRL1
SMARCA4
TICAM1
AAMP
TNFRSF19
SHFM1
GNAI1
CCNT1
ILF3
PAK4
TRAF3
PSME3
RARA
PIAS3
RAF1
TRIM5
ANKRD17
AP3B1
RFXAP

KRAS
IRAK2
SORT1
TRIM45
SAA2
IL6ST
ELP2
TPP2
UBQLN1
TRAFD1
MAP3K7
CAPRIN1
GDF11
CDK4
ACVR1B
SEMA4F
RNASEL
DHX9
COX5B
PLXNC1
IREB2
CALCOCO2
TANK
GATA4
VPS45
KIAA0368
DAB2IP
RIPK1
SMAD6

ADAM10
ITGAV
SLC40A1
PPP3CA
SCAF11
SLC15A4
LMBR1L
RB1
CLTC
SMAD4
SOD1
AKT1
BCL10
ISG20L2
HDGF
SNX27
RFX5
IL17RD
NFKBIZ
KIAA0226
TIFA
PDGFC
SKP2
PIK3R1
ATG12
G3BP1
IRAK1BP1
EGFR
VLDLR

NR6A1
PARD3
TCF7L2
ITGB1
TIRAP
CRIM1
THRB
EIF4E
EPS8
CCL28
PDK1
MERTK
TOMM70A
PRKCA
CNOT8
PPARGC1B
DCK
NODAL
BRAF
NCK1
FANCC
ADIPOR1
UBR1
VAV2
SHC1
NLRX1
ADAR
RUSC1
JAK1

RBP7
MAPKAPK2
REL
AZI2
PSMD6
SENP2
CASP3
ERAP1
RICTOR
LEAP2
HNF4G
RAD21
ABCA1
ARF6
TSC1
DDX21
CASP7
HIF1AN
C2
ILK
TRIM66
STAT6
SMAD3
STIM1
CTNNB1
FADD
LTBP3
NKIRAS2
IRF2

STAT3
SEMA4C
TET2
MFF
MAP2K1
PTK2
PCBP1
LIMS1
ROBO1
TRIM56
SEMA3E
SOCS6
MFN1
PRKCE
SOCS5
TRIM8
PTEN
MALT1
CEBPB
MYD88
RELA
SAA1
PSMD1
STAT5B
NR1D2
BTC
PDE12
DHX36
TRAF6

PSMD2
INHBC
NR2F1
IP6K1
CRLF3
C8G
ANO6
ULK1
NR2C2
JUN
IL17RA
GRB2
UBE2N
ERN1
RAD23A
PTPN11
ELMOD2
ZFPM1
PRKRA
PAK2
NRIP1
RNF41
MTA1
TBK1
IRAK1
SEMA4B
PURA
NR2F2
SP1

PTCH1
PPP1CC
MAPT
EPOR
PTMA
HSPA14
USP7
PLSCR1
QRF1
SLX4
LRRK2
HMGB1
IL1RAP
SIAH1
PLXNB2
HDAC2
FAM3C
GMFB
IGF2R
PCBP2
PELI1
IRAK4
5-Mar
COPS8
MTOR
HMGN2
MAP3K3
SREBF2
DAXX

BMP2
RXRB
SKIV2L
EHMT2
HSPA1L
APOM
MICB
TRIM27
TRIM13
LGR4
MIR499A
NRAS
CHUK
LTBR2
DDX3X
PPP3R1
CEBPD
ORM2
ORM1
TRIM26
CNTF
TICAM2
CFB
YJFN3
PRKDC
HP
ITGB3
FIGL2
OTUD7B

Brown

IKBKG
SRXN1
CFH
MASP2
PSMC4
BID
C8B
FAS
GRN
USP2
C6
RFXANK
RORA
JMJD6
WDR62
PSMC5
F11
BIRC5
ESR1
TYRO3
PSMD8
PSMA7
TRIB3
E2F1
RNF125
PQBP1
TRADD
MAPK3
RIPK2

NDRG1
NFKBIB
RPS19
PIK3R2
GSK3A
TFR2
C5
RPL19
C1QBP
YWHAE
PPARGC1A
ZBTB16
MDK
AIP
GHR
KNG1
HRG
HDAC1
LEPR
ARG1
WDR34
DUSP1
SOCS2
COPS5
CAT
ACO1
PKN1
PACSIN1
C3

RBCK1
IRF3
PRMT1
MAP2K2
TRAF2
PIN1
ADM2
INS-IGF2
EPO
GDF15
ADRM1
TRIM28
NR1H2
HSPBP1
CNPY3
RBP4
TRIM65
HSPD1
NR1I2
CCNA2
TSLP
CDKN2A
PTGES2
SERPING1
PPP4C
TCEB1
MOV10
C8A
TSC22D3

NLRP14

XDH

IFNGR2

RCAN1

PSMD4

C1R

SPON2

S100A1

AZGP1

FGFR4

UBXN1

HDAC11

ALB

PRKCD

AIMP1

HMGB2

KLKB1

MBL2

PSMC3

AVPR1A

PLK1

OTUB1

CD320

MLST8

FABP4

CD14

IRF2BP1

HSPA4

FGA

GNB2
PPP1CA
HRAS
DDIT3
IMPDH2
SHARPIN
SSTR2
C1S
DDX41
TRAIP
PROS1
HSF1
PSMD13
DEFB132
INSIG1
PPIA
XRCC6
SRC
GUCA2A
NMB
S100A10
LPA
SCAMP5
ATG9A
GNB2L1
GCGR
CKLF
MIF
EGLN2

Green

ORAI1
CCL14
PLXND1
SEMA3G
KAL1
DCN
VIM
FLT4
TNC
LTBP1
ELN
PTGER3
NGFR
CALCRL
GLP2R
TIE1
PTGS2
SCARF1
SEMA3C
RARB
FGFR1
EDN1
NOX4
LTBP4
CD200
SLC22A17
PDGFB
PROCR
JAG1

SMAD7
BMX
FLT1
CLEC11A
PTN
CAV1
NOD1
OGN
ENG
CXCL12
GLI1
BMP5
C7
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ANGPTL1
AKT3
NRP2
CTGF
LTBP2
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TEK
EGR1
INHBA
NR4A1
F2RL3
KDR
DLL4
WNT2B
NGF

APLNR
PDGFRA
ETS1
ANXA1
EDNRB
ANGPTL2
NOV
IL33
CCL21
THBS1
SEMA6D
BMPR1B
ACVRL1
MAP3K12
PELI2
FGF7
FURIN
GATA6
CYR61
TINAGL1
CTSK
SLIT2
NOTCH1
VEGFC
ITPR1
EDNRA
FLI1
CYSLTR2
GJA1

RASGRP3

ANGPT1

FGF18

GDF6

IL34

STC1

CSRP1

PTH1R

FSTL1

TGFBR2

PTX3

EDIL3

DEFB1

JAM3

HDGFRP3

AXL

SEMA6B

NXN

BDKRB2

NPR1

ZFPM2

FOS

FSHR

PDGFD

S1PR1

BCL2

SCG2

RARG

F2R

Grey

BGN
GAS6
GDF3
TMEM173
JAG2
NTF3
APOD
TCF4
PDGFA
GRK5
FAM19A5
LCN6
CALCR
MPO
CRLF1
TNFRSF12A
NOS2
SELE
NOX1
PGLYRP1
SEMA3B
NR1H4
IGF1
NLRP2
NR1H3
IFNGR1
HMGB3
MAP2K3
CDH1

TG
ADRB1
HSPA5
IL17RB
OGFR
NTN1
FGFR2
TNFRSF1A
GAL
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FGF10
FGF22
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PRLH
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SREBF1
SCARB1
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NOX3
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PAK3
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HOXA9
ADCYAP1R1
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TOLLIP
CEACAM1
CXCL2

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OPRK1
ACHE
CETP
LPCAT2
SH3BP2
DEFB127
OAS1
NOS1
THPO
CMA1
PSME1
SEMA4G
BLNK
PGC
NCR2
MLN
POLR2F
CTSG
TRIM9
NFKBIA
PSME2
R3HDML
AVP
ANGPT4
PDYN
PAK7

OXT
BPI
CST4
SYP
FGF14
MLNR
PSMD7
STUB1
CTSH
CGB2
CGB
FCGRT
AMH
CLEC4M
GPI
FGF21
HAMP
NAMPT
MET
VIPR2
CRHR2
GHRHR
RLN1
DKK1
CSF3
CCL1
DHX58
PPY
GNRHR

CD81
CALCA
IL23A
ING4
OPRM1
IL17A
IL17F
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NR2E1
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LIFR
C9
SELK
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RBP1
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CISH
VIPR1
POMC
GCG
TACR1
REG1A
IL1R2

IL1RL2
IL1RL1
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BMP8B
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GHRH
FGF23
ESRRB
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IFIT2
INSL6
INSL4
IFNA6
TNFSF11
NFYB
IAPP
GHSR
TNFSF10
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OBP2A
NPY
CNTFR
SFTPA1
LRP1
ACVR1C
C4BPA
MC3R

SLPI
SDC4
SEMG2
EDN3
SEMG1
IL17C
CEACAM8
EREG
SLC10A2
AMELX
PSPN
TNFSF14
GNRH2
FOXA2
BMP2
DEFB129
GDF5
SLURP1
NR1D1
HTN1
IFI6
AVPR2
EDN2
WFIKKN1
MCHR1
GALR3
FGF13
SHC2
LBP

ECSIT
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BST2
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GFAP
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FSHB
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PPARG
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RXFP2
TPT1
KL
NTS
SFTPD
SPINK5
TSHB
RSAD2
GRP
HRH4
MTNR1B
OASL
CD36

CGA
LACRT
NMBR
TEC
AGT
STAB2
GH2
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IL1F10
CDK9
IL11RA
IFNA21
PI15
LHCGR
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FGF2
EGF
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INHBE
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IFITM3
IFNAR1

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CBLC
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FCN3
RXRG
SEMA6C
DUSP10
S100A7
ACTA1
LEFTY2
GDF7
REG3G
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MUC4
FABP2
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IFNK
SHC3
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DAK
CTF1
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PTH
TRIM36
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BMP3
BMP6
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SEMA3D
ROBO3
WNT3A
TMSB4Y
PIK3AP1
TRIM42
ADCY8
RAET1L
NRG1
TRIM63
COLEC12
NRG2
CD1D

FGF17
WNT9B
FBXW5
GIP
AMFR
AGRP
NR2F6
FCN2
IL6R
PTGER1
SQSTM1
PGLYRP2
SCGB3A1
PSMC2
DCD
CXCL16
FGF19
IL23R
TRIM58
SLAMF9
FCGR3B
ATF3
IL20
INHBB
BMP10
S100A12
NPPC
FAM19A4
RETNLB

IL17RE
IL17RC
PPBP
PF4
UCN
S100P
PGRMC2
CAMP
PLXNB1
ESM1
ERAP2
TLR3
IL3
GDF9
IL31RA
DEFA5
DEFA4
SPAG11B
NFIL3
FIGF
AQP3
SUGT1
CLEC1B
RNASE7
IFI27
IL25
CMTM5
RAG1
CRABP1

TK2
HSP90B1
HTR3A
CHP2
PDIA3
IGF2
KLK1
ANGPTL4
PRDX2
MTNR1A
BMP1
GDNF
WFDC12
TRIM49
UBE2V2
IL13
NR0B1
IL1RAPL1
GP2
SDC2
RAC3
NRG4
CMTM8
STAT2
TRH
OBP2B
INSR
NRTN
RLN3

APLN
NLRP5
RXFP1
LGALS4
ANGPTL7
PRL
AZU1
INSL5
IL17D
ESRRA
GLRX
TNFRSF10D
NLRP13
CCR9
RGMB
PELI3
IL20RB
LEP
NLRP6
NPPA
PCSK1
CALCB
A2M
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LCN15

STAP2
HTR3C
SPAG11A
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CALR
GPHB5
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APOBEC3B
NLRP11
DEFB124
DEFB123
DEFB119
AGTR2
DEFB112
GREM2
AEN
ADIPOQ
PENK
DDX60L
SIAH2
NLRP10
EPGN
RXFP3
GALR2
TCHHL1
TRIM61
CCR3

FAM19A1
MRGPRX2
FAM3B
DEFB108B
S100A7A
COLEC10
CCR10
LCN12
SIVA1
IFNE
ROBO2
SIGIRR
IFITM2
MC2R
SFTPA2
PARK2
IRF7
DLK1
PDIA2
SHC4
IFIT1
NLRP9
DEFB128
HTR3E
HTR3D
DEFB131
RXRA
DEFB106A
DEFB105B

IFNA10
FGF3
PPARA
TMPRSS6
CCK
ANGPTL5
ISG15
DMBT1
LCN10
PLA2G2A
IFNA2
PPP3R2
OSTN
NOXA1
CGB5
CXCL17
SERPINA3
PRTN3
FGF16
ESRRG
SIGLEC15
TPSB2
S100A7L2
FABP12
MAP3K5
GALP
ELANE
NKIRAS1
IFNA1

PLCG2
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FAM3D
CALM1
FAM19A2
RORB
MIR107
MIRLET7C
MIR373
LCN8
MAFB
AGER
RNF5
NEU1
HSPA1B
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CSHL1
MICA
DEFB121
TRIM15
IL31
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TECPR1
HTN3
CRLF2
DEFB136
TRIM71
MIR23B

MIR181A2
MIR141
MIR200C
MIR517A
MIR10B
MIR187
MIR517C
MIR125B2
MIR328
MIR125B1
MIR23A
MIR485
IGLV5-52
IGLJ4
IGLJ5
IGLJ7
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TRGJP1
TRGV9
TRAV7
TRAV8-7
TRDV2
TRDJ4
TRDJ2
TRDJ3
TRAJ61
TRAJ59

TRAJ58

TRAJ57

TRAJ56

TRAJ54

TRAJ53

TRAJ52

TRAJ50

TRAJ49

TRAJ47

TRAJ46

TRAJ45

TRAJ43

TRAJ41

TRAJ40

TRAJ35

TRAJ33

TRAJ30

TRAJ29

TRAJ28

TRAJ27

TRAJ26

TRAJ23

TRAJ22

TRAJ21

TRAJ20

TRAJ19

TRAJ14

TRAJ13

TRAJ11

TRAJ10
TRAJ9
TRAJ8
TRAJ7
IGHE
IGHD1-26
IGHD5-24
IGHD2-21
IGHD6-19
IGHD5-18
IGHD5-12
IGHD2-8
DEFB117
CGB8
LTB4R
CCL27
DEFB133
GPR33
MUC5AC
DEFB116
DEFB115
MIR208B
PLXNA4
C4B
IGHD1-14
IFNA14
IGHD4-11
IGHD4-4
IFNA13

IFNA17
IGHD1-1
IGHD7-27
IGHD1-20
TLR9
DEFA3
TDGF1
CORT
KIR3DL3
IL10RB
C4A
TRBV6-8
INS
NOX5
CARD18
PDF
RN7SL1
MC1R
TRAV30
GH1
MIA
GDF2
MIR3148
TXNIP
GDF10
MIR212
VTRNA2-1
CCL23
TRBV16

Red

CCL15
TRAJ36
MIR29A
NR2E3
TRAJ37
TAC1
CHGB
CHGA
BMP7
FGF9
FGF8
FGF6
OPRD1
PRLHR
NPPB
DEFB126
PCSK2
PRKCG
VGF
PYY
AMHR2
CSH1
NR5A1
PROK1
IL9
CRH
GPHA2
MCHR2
SST

	TMSB15A
	TMSB15B
	AIRE
	NLRP4
	FGF11
	PROK2
	FABP7
	DEFA6
	FREM1
	LALBA
	NLRP7
	GKN1
	S100G
	TRHR
	IFNW1
	HTR1A
	UCN3
	SCN5A
	SLIT1
	AVPR1B
	MIR130A
	TRAJ48
	CSH2
	IFNA4
Turquoise	FGR
	CASP10
	ITGAL
	CX3CL1
	CD79B

CD4
BTK
FYN
PLAUR
TYROBP
CD22
LTF
WAS
SLC11A1
MARCO
CD74
HGF
RUNX3
BIRC3
TYMP
SLAMF7
BTN3A1
TNFRSF1B
MSR1
LCP2
TNFRSF17
UTS2
TNFRSF9
TNIP3
CYBA
TRAF1
PRKCQ
SPI1
BCL3

FCGR2B
ACAP1
CD5L
TXK
SEMA3A
ARHGAP15
IL4R
PTPRC
GSTP1
FCN1
SIGLEC1
ICAM1
IRAK3
LYZ
FLT3LG
CD209
NLRC4
IL5RA
NLRP1
CEBPE
CECR1
TREM2
IL12RB1
OSM
LGALS2
HMOX1
APOBEC3H
GRAP2
CSF2RB

IL2RB
GZMB
LGMN
PLTP
MMP9
CD40
HCK
SAMHD1
WFDC2
TLR8
CD40LG
TNFSF13B
OLFM4
CCL22
CCL17
MEFV
IL21R
CSK
IL7
CD37
RETN
IL27RA
EBI3
SIGLEC8
CD79A
SIGLEC6
SIGLEC5
JAK3
PIK3CG

CCL24
SERPINE1
TRIM14
AKNA
TNFSF8
RLN2
DDX58
PTGDS
ICAM2
CCL7
CCL2
CCL8
PF4V1
AREG
IL2
HSPA8
IL10RA
OAS3
OAS2
VDR
IL26
IFNG
PTPN6
AICDA
BTN3A3
TRIM38
CCR6
LY86
HBEGF

ITK
IL12B
FGF1
STC2
CD86
GNAI2
IL1A
ZAP70
CYTIP
IFIH1
STAT1
GNLY
IL18R1
IL18RAP
SDC1
PLA2G4A
RGS2
SLAMF1
CD48
CR2
PIK3R3
FASLG
SPP1
NR4A3
CSF3R
CD274
TNFSF18
PTK2B
PPP3CC

TNFRSF8

TRIM6

FABP3

CCRL2

CXCR4

FLT3

CD244

LY9

PLAU

CD97

OPTN

NCKAP1L

IL13RA2

ZBP1

IL9R

NDP

TREM1

CXCL6

IRF1

PTGER2

KIR2DL1

OPRL1

IL1B

TNFSF9

CD70

FFAR2

HCST

CCR7

STAT5A

CFP
HSPA2
IL22
IL17B
RAC2
APOBEC3A
APOBEC3F
SIGLEC9
RIPK3
GMFG
ULBP3
NFATC1
IDO1
RFTN1
LGALS3
TRIM21
TRIM22
CARD6
CRP
CHIT1
RARRES3
CD180
IL2RA
IL15RA
KLRD1
KLRC1
HAVCR2
IL6
IL10

TLR4
CD72
IRF4
BPHL
TLR2
ARRB1
MMP7
CASP1
STAT4
IL21
CXCL9
CD27
CELA1
ESR2
PSTPIP1
BCL2A1
PML
ITGAX
NLRC5
CMTM2
IRF8
ARRB2
PIK3R5
SECTM1
PMAIP1
VAV1
TRPM2
NLRP12
SIGLEC10

SYTL1
CD53
XCL1
XCL2
FCGR2A
RORC
S100A8
SNCA
OSMR
CXCL14
DOK3
IL2RG
FCGR1A
IL18
VENTX
MR1
CMTM7
CD8A
LY96
CXCL13
GHRL
MX1
TNFRSF14
CD1A
CD1C
CD1B
CD1E
SLAMF8
FCER1G

IFNAR2
C1QC
TNFRSF13C
ABCG1
ITGB2
S100B
TOR2A
CD3G
CXCR5
CCR5
SIGLEC11
TNFSF13
SDC3
GBP2
VCAM1
NLRP3
SLAMF6
IL24
CTSS
TNFAIP8L2
PGLYRP4
S100A9
CXCR1
TRAT1
IFI16
AIM2
CTLA4
ICOS
CXCL3

CXCL5
CXCL1
CCR1
ZC3H12A
IL15
F2RL2
F2RL1
CSF2
MB21D1
IL22RA2
RAET1E
FABP5
CTSB
TNFRSF11B
SYK
CYBB
TSHR
PRKCB
CLEC4E
CLEC4D
MC4R
B2M
GREM1
NOD2
CD3D
LAIR1
KIR3DL1
CD300A
NLRC3

PNOC
PTGDR
CX3CR1
TAP1
IL7R
IL12A
INPP5D
LGALS9
SIGLEC7
CXCL10
CXCL11
RNASE2
RNASE3
PTAFR
ITGAM
TPST1
FABP6
SERPINB9
FPR2
FPR1
PTGER4
PIK3CD
S100Z
IFNB1
C3AR1
CD8B
CCL11
ISG20
CXCR6

CLEC7A
IL16
RASGRP1
CCL19
HSPA6
CYSLTR1
C1QB
C1QA
OLR1
XCR1
TLR10
TLR1
TLR6
CMKLR1
P2RY14
DES
UCP2
BDNF
CD19
CD28
HLA-DQB1
CIITA
CCR8
PRF1
CCL13
RNF135
CSF1R
LCK
PMCH

MX2
KLRC4
CCR4
SH2D1A
CSF1
SOCS3
FAM19A3
IL3RA
SOCS1
IFITM1
CD300LF
CD300E
BTN3A2
CXCR3
TNFRSF4
TNFRSF18
TRPV2
SEMA4D
CARD9
PDCD1
KIR2DL4
NCR1
HLA-DRB1
SEMA4A
S100A5
TLR7
HLA-DQA1
CASP4
C5AR1

GZMM
PDCD1LG2
MAP1LC3C
CLEC9A
SIRPA
CLEC4C
CSF2RA
SLC29A3
CARD11
HLA-DRB5
SH2D1B
CD247
CD3E
RAET1G
FCGR3A
HLA-DOA
HLA-DMA
PSMB8
TAP2
HLA-DRA
CARD16
NCR3
LST1
HLA-C
LILRB3
HLA-E
HLA-G
HLA-F
TMSB4X

KLRC2
KLRC3
CLEC6A
HLA-H
HLA-A
IGKC
IGKJ5
IGKV4-1
IGKV5-2
IGKV6-21
IGKV3D-20
IGKV6D-41
IGKV3D-11
IGKV1D-42
IGLV4-69
IGLV8-61
IGLV4-60
IGLV6-57
IGLV11-55
IGLV10-54
IGLV1-51
IGLV1-50
IGLV5-48
IGLV1-47
IGLV7-46
IGLV5-45
IGLV1-44
IGLV7-43
IGLV1-40

IGLV5-37

IGLV1-36

IGLV2-33

IGLV3-32

IGLV3-27

IGLV3-25

IGLV2-23

IGLV3-22

IGLV3-21

IGLV3-19

IGLV2-18

IGLV3-16

IGLV2-14

IGLV3-12

IGLV2-11

IGLV3-10

IGLV3-9

IGLV4-3

IGLV3-1

IGLJ1

IGLJ2

IGLC2

IGLJ3

IGLC3

IGLJ6

IGLC7

TRGC1

TRGV5

TRGV4

TRGV3
TRBV6-1
TRBV4-1
TRBV6-4
TRBV7-3
TRBV9
TRBV10-1
TRBV11-1
TRBV6-5
TRBV6-6
TRBV5-5
TRBV7-6
TRBV5-6
TRBV5-7
TRBV5-1
TRBV4-2
TRBV19
TRBV20-1
TRBV24-1
TRBV25-1
TRBV27
TRBV28
TRBJ2-1
TRBJ2-2
TRBJ2-3
TRBJ2-4
TRBJ2-7
TRBC2
TRAV2

TRAV3
TRAV4
TRAV5
TRAV6
TRAV8-1
TRAV9-1
TRAV10
TRAV12-1
TRAV8-2
TRAV8-3
TRAV13-1
TRAV12-2
TRAV8-4
TRAV13-2
TRAV14DV4
TRAV9-2
TRAV12-3
TRAV8-6
TRAV16
TRAV17
TRAV18
TRAV19
TRAV20
TRAV21
TRAV22
TRAV23DV6
TRDV1
TRAV24
TRAV25

TRAV26-1
TRAV27
TRAV29DV5
TRAV26-2
TRAV34
TRAV35
TRAV36DV7
TRAV38-1
TRAV39
TRAV40
TRAV41
TRDC
TRAJ44
TRAJ42
TRAJ39
TRAJ38
TRAJ34
TRAJ31
TRAJ25
TRAJ24
TRAJ18
TRAJ17
TRAJ16
TRAJ12
TRAJ6
TRAJ5
TRAJ4
TRAJ3
TRAJ2

TRAJ1
IGHA2
IGHG4
IGHG2
IGHA1
IGHG1
IGHG3
IGHD
IGHM
IGHJ2
IGHJ1
IGHD3-22
IGHD3-16
IGHD2-15
IGHD6-13
IGHD3-10
IGHD3-9
IGHD3-3
IGHD2-2
IGHV6-1
IGHV1-2
IGHV1-3
IGHV2-5
IGHV3-7
IGHV3-11
IGHV3-13
IGHV3-15
IGHV3-16
IGHV1-18

IGHV3-20

IGHV3-21

IGHV3-23

IGHV1-24

IGHV2-26

IGHV4-28

IGHV3-33

IGHV4-34

IGHV3-35

IGHV3-38

IGHV4-39

IGHV1-45

IGHV1-46

IGHV3-48

IGHV3-49

IGHV5-51

IGHV3-53

IGHV1-58

IGHV4-61

IGHV3-66

IGHV1-69

IGHV2-70

IGHV3-73

IGHV7-81

LAT

KLRK1

UBD

IRF9

TNFRSF25

IFI30
KIR2DS4
IGLC6
IGLV9-49
IGHV3-64
HLA-DPB1
IGKV3D-15
IGHV4-59
IGHV3-74
IGKV6D-21
IGHV3-72
IGHD6-25
TRBV2
LTA
TRGC2
IGHD4-23
LTB
IGHD4-17
IGHD6-6
IGKV3D-7
TRBV10-2
TRBV5-4
HLA-DPA1
IGHV4-31
TAPBP
IGHV3-43
TNF
TRBV29-1
TRGV2

HLA-B
TRBV30
HLA-DQA2
TRBV3-1
IGKV2D-30
TNFSF12
APOBEC3G
IGKV1D-8
IGKV1-6
IGKV1-37
IGKV3-20
LILRA4
IGKV1D-33
LILRA2
IGKV1-17
KIR3DL2
TNFRSF13B
IGKV1-8
IGKV1-16
HLA-DOB
IGKV1D-16
IGKV2-24
IGKV3-11
IGKV2D-24
TRBV11-2
IGKV1-9
IGKV1-33
IGKV1-39
IGKV2D-28

HLA-DMB
IGKV1D-43
IGKV1D-17
IGHJ3
IGKV3-7
IGKV2-30
IGKV2D-29
IGKV1-12
IGKV1-5
LEFTY1
KIR2DL3
IGKV2-28
IGKV3-15
APOBEC3C
IGKV1-27
INSL3
NAIP
IGKV1D-37
IGKV2D-40
IGKV1D-39
TRBV6-7
TRBV7-7
TRBV7-4
IGKV1-13
LYN
TRAV1-1
TRAV1-2
TRDV3
MMP12

	S1PR2
	IGHV3-30
	CCL5
	IGKV2-40
	TRBV12-3
	TRBV12-5
	CCL4
	CCL18
	TRBV14
	TRBV10-3
	CCL4L1
	CCL3L3
	TRBV13
	TRBV18
	IGKV1D-13
	TRBV11-3
	IGHV4-4
	TRBV15
	TRBV12-4
	CCL3
	TRAC
	TRBV17
	TRBV7-9
	IGLV2-8
	IGKV1D-12
Yellow	CFTR
	ITGA3
	CCL26
	GIPR

IL20RA
TMSB10
KCNH2
NCK2
NFKB2
SCTR
TRAF5
SEMA5B
PTHLH
APOH
SEMA6A
IL11
ITPR3
SERPIND1
LGALS1
NFATC4
HNF4A
ELF4
PYCARD
AQP9
RAB11A
PDGFRL
LHB
RELB
NUMBL
TGFB1
UNC93B1
LTBR
CCL20

PROC
ARTN
TRIM67
PGF
CRHR1
KCNJ8
PLG
INHA
PI3
UNC5CL
VASP
MASP1
ATF4
LIF
IRF5
RHBDF2
PLXNA3
TRIM47
ANGPTL3
VAV3
CD63
RAC1
PAK6
DUOX1
CASP6
RBP5
DUOX2
IGF1R
CMTM3

TNFRSF11A
NR1I3
CRABP2
PKLR
AHSG
LECT2
TNFRSF21
NFKBIE
TRIM50
ZC3HAV1L
GNRH1
PTGES
PAK1
UCHL1
FZD1
PGLYRP3
ZYG
LCN1
GPSM1
SSTR5
RPS6KA4
S100A11
TGFA
MST1R
HFE2
AR
ADRB2
NUDT6
ADRBK1

VEGFB
TNFRSF10C
TNK1
PTPN2
CLCF1
SPHK1
RABEP2
PLEC
OXTR
TNFSF15
ANXA2
BMP8A
CMTM4
PRKX
ACTG1
GAST
MUC1
NRG3
TLR5
S100A3
S100A16
CLDN4
S100A13
S100A14
CTSE
CGB7
S100A2
ANXA4
ANXA6

IL27
S100A6
TUBB3
TPM2
PLXNB3
TBKBP1
DEFB134
LCNL1
NTF4
IKBKE
CCL16

Supplementary Table 3. Proportion of variance explained by each principal component

PC	Freq
PC1	0.928236
PC2	0.046466
PC3	0.007112
PC4	0.005018
PC5	0.002986
PC6	0.001962
PC7	0.001696
PC8	0.001324
PC9	0.001022
PC10	0.000986
PC11	0.000859
PC12	0.0007
PC13	0.000561
PC14	0.00038

PC15	0.000277
PC16	0.000243
PC17	0.000172
