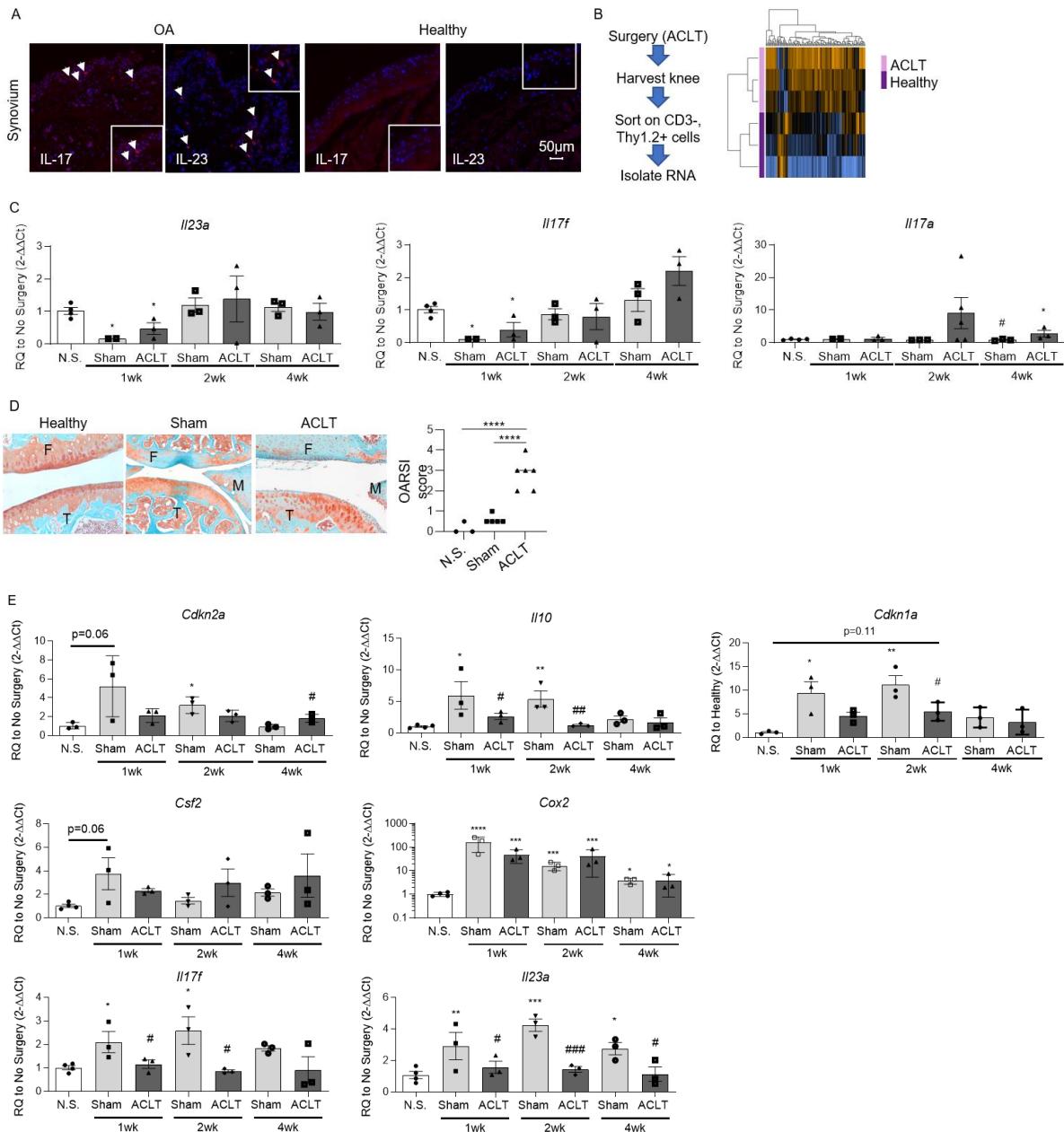
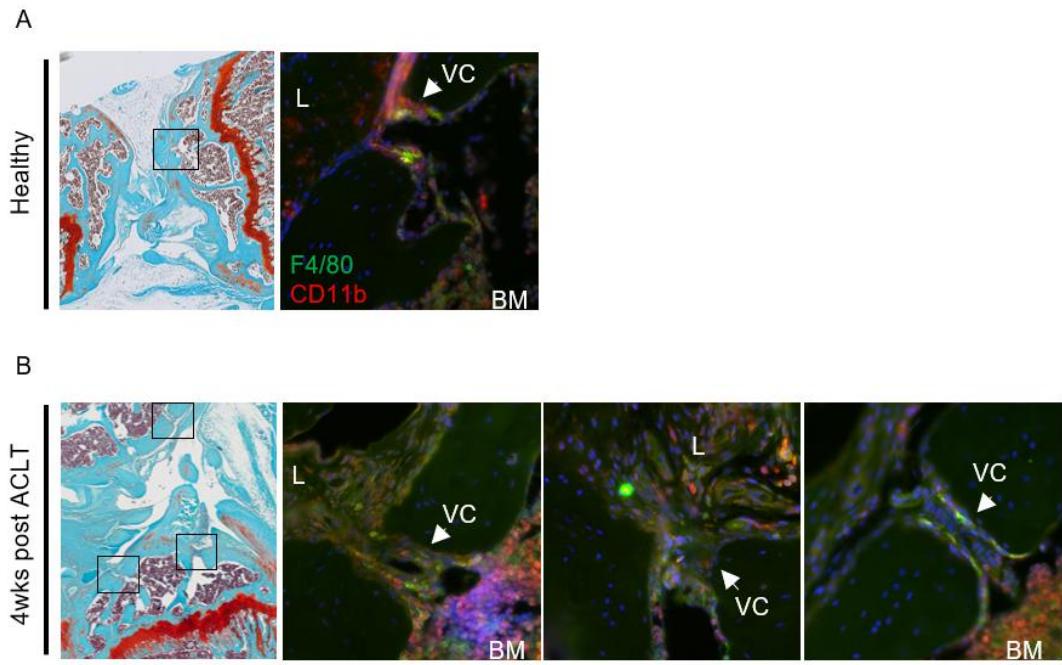


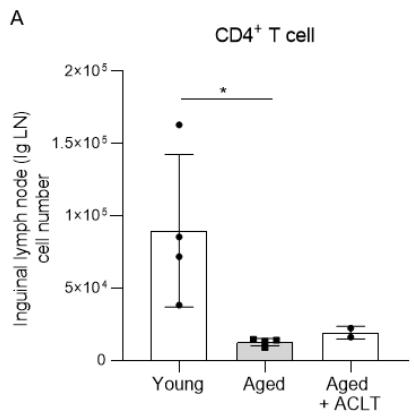
Supplemental Figure 1. No lymphoid cell number changes are observed in the joint 2-and-4 weeks post ACLT. Numbers of IFNy+ CD4+ T cells in the joint are similar to no surgery at 1 week (A), 2 weeks (B), and 4 weeks (C). (D) is an example dot plot from an ACLT joint 2 weeks post surgery demonstrating the low number of IL-4+ cells (30) for quantitation. This sample is representative of most joint samples for IL-4 levels. Lymphoid cell numbers in the joint are similar to no surgery at 2 weeks (E) and 4 weeks (F) post surgery.



Supplemental Figure 2. (A) Immunofluorescence of human healthy and OA synovium. Only OA tissue exhibits IL-17 and IL-23 staining. Joint and inguinal lymph node post injury gene expression, flow cytometry, and immunohistochemistry characterization. (B) Nanostring pathway analysis of ILCs sorted from 2wk post ACLT vs no surgery joints. (C) Inguinal lymph node PCR from young mice with sham or ACLT 1, 2, and 4 wks post surgery (n=3). Stars represent significance compared to the no surgery group, pound signs indicate significance compared to opposite surgical group, and when p is indicated numerically, groups being compared are specified with a line. Groups are only statistically compared within each time point. (D) OARSI scores and representative images of sham joints 4wks post surgery. E. Whole joint PCR from young mice with sham or ACLT 1, 2, and 4 wks post surgery (n=3). Same statistical scheme as used in C. * $p<0.05$, ** $p<0.01$, *** $p<0.001$ and **** $p<0.0001$.

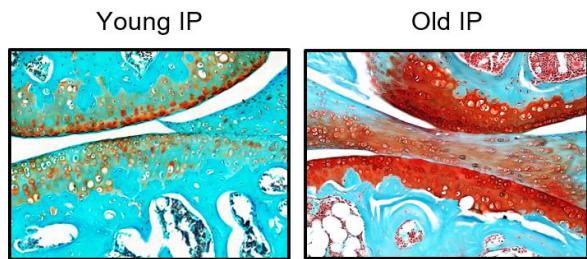


Supplemental Figure 3. Vascular channels change with joint trauma. Joints were sectioned and regions with ligaments insertions were imaged to identify vascular channels. Vascular channel=VC. Bone marrow=BM. Ligament=L. In healthy joints (A) $F4/80^+ CD11b^+$ cells are identified lining the walls of the vascular channels. In mice that have undergone ACLT (B) $F4/80^+ CD11b^+$ cells traffic through these vascular channels. More vascular channels were observed in ACLT joints than in healthy joints.

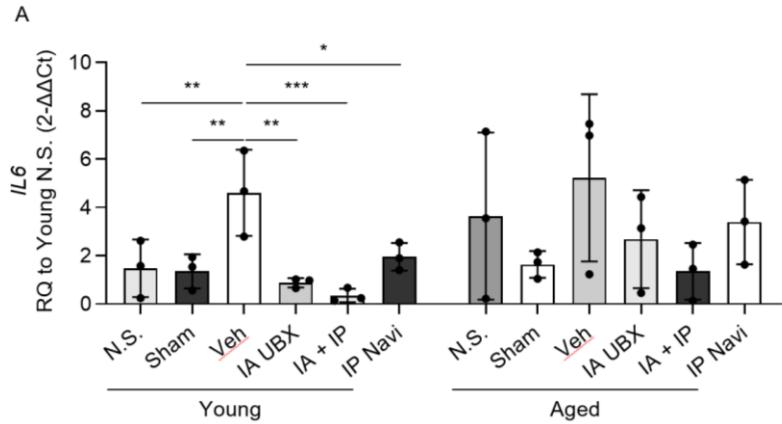


Supplemental Figure 4. Aging decreases lymph node lymphoid cell number. (A) Inguinal lymph node total CD4⁺ T cell number in aged vs young mice. * $p<0.05$.

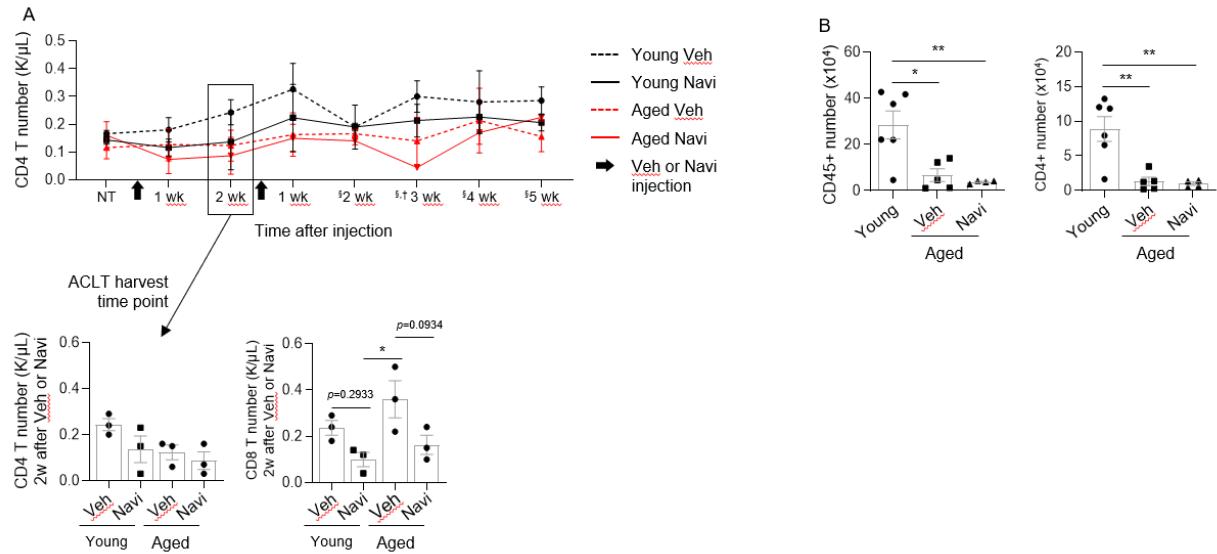
A



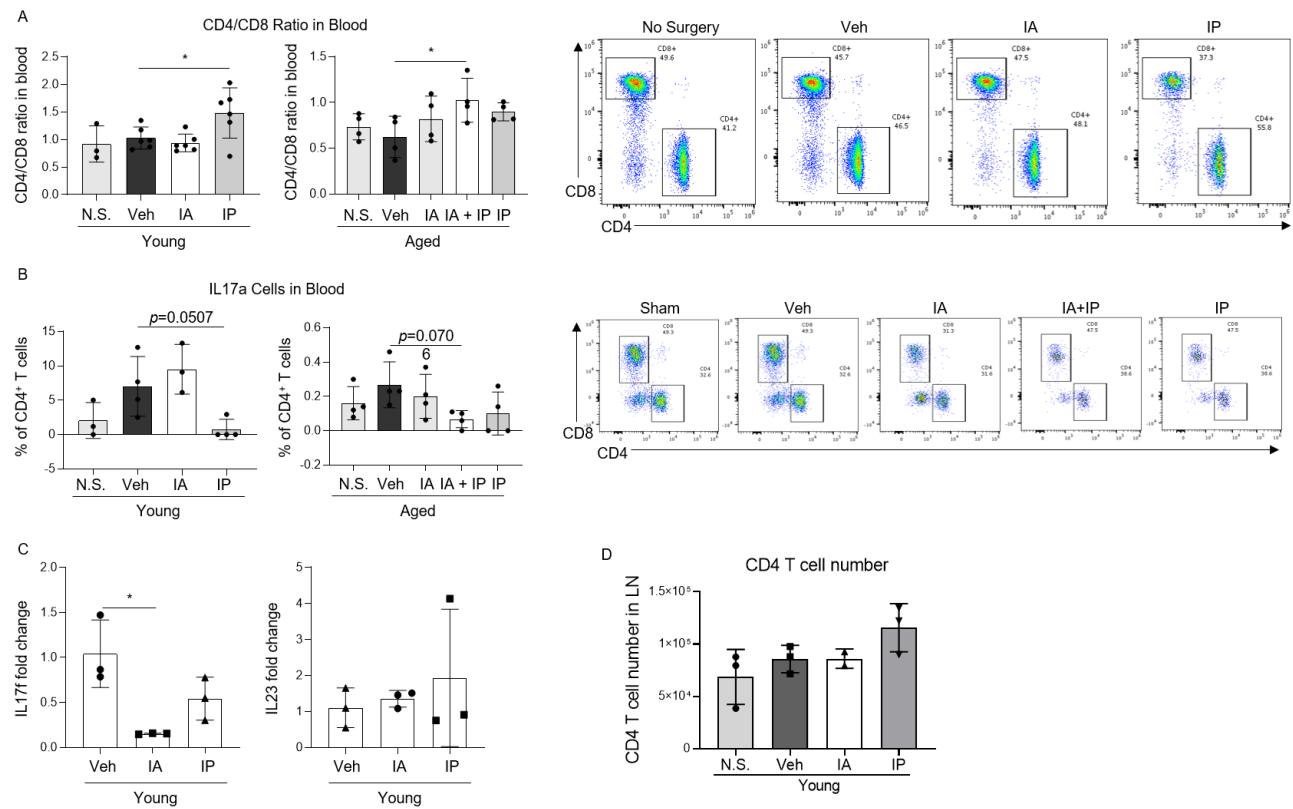
Supplemental Figure 5. Senolytic treatment of senescent cells enhances repair and decreases Th17 signature. (A) Representative histological images of systemic Navitoclax treatment in young and aged animal increase the cartilage repair.



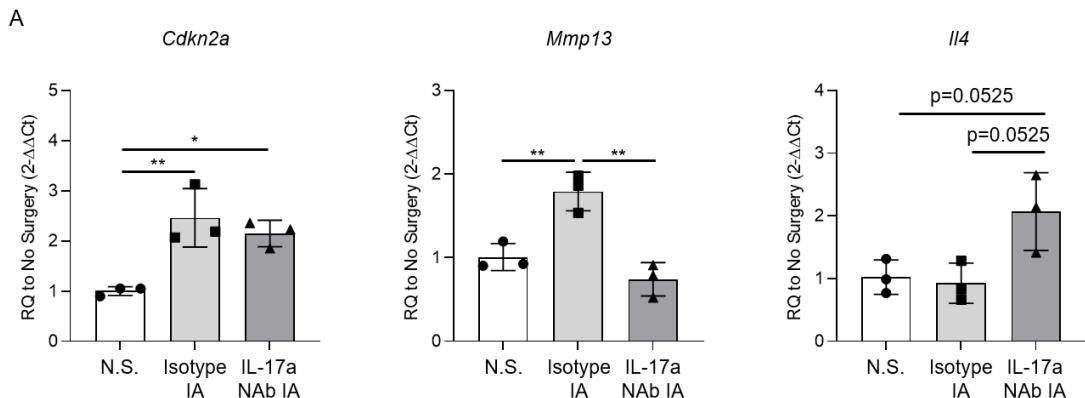
Supplemental Figure 6. SASP factor IL6 expression. (A) Quantification of mRNA expression of IL6 in articular joints before and after senolytic treatment in young and aged mice. * $p<0.05$, ** $p<0.01$, and *** $p<0.001$. Experimental groups are compared to the control group (Veh).



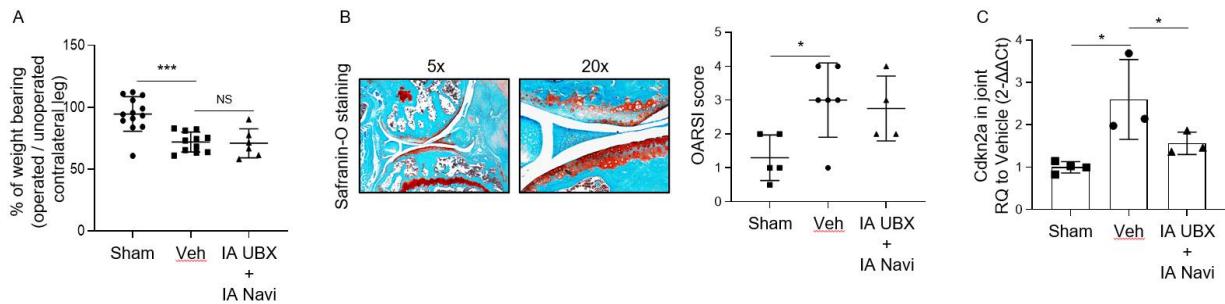
Supplemental Figure 7. Effect of senolytic treatment on lymphoid cells. (A) Effect of Navitoclax on blood T cell numbers after Vehicle or Navi treatment. N=2 for Young Veh and Aged Navi for the time points marked with § . N=2 for Aged Veh for the time point marked with †. N=3 for the rest of time points for all groups. * $p<0.05$. (B) Effect of Navitoclax on inguinal lymph nodal CD45 and CD4 cells 2 weeks after treatment. * $p<0.05$, and ** $p <0.01$.



Supplemental Figure 8. Systemic senolytic treatment in ACL transected mice induces systemic immune changes. (A) CD4/CD8 ratio change in young and aged mice after treatments, and corresponding flow plots. (B) IL17a cell percentage in blood, and corresponding flow plots. (C) IL17f and IL23 fold change in young animal inguinal lymph nodes. * $p<0.05$. Experimental groups are compared to the control group (Veh). (D) Quantified CD4+T cell numbers from inguinal lymph nodes of young mice; n=2-3 for each group. * $p<0.05$. *** $p<0.001$.



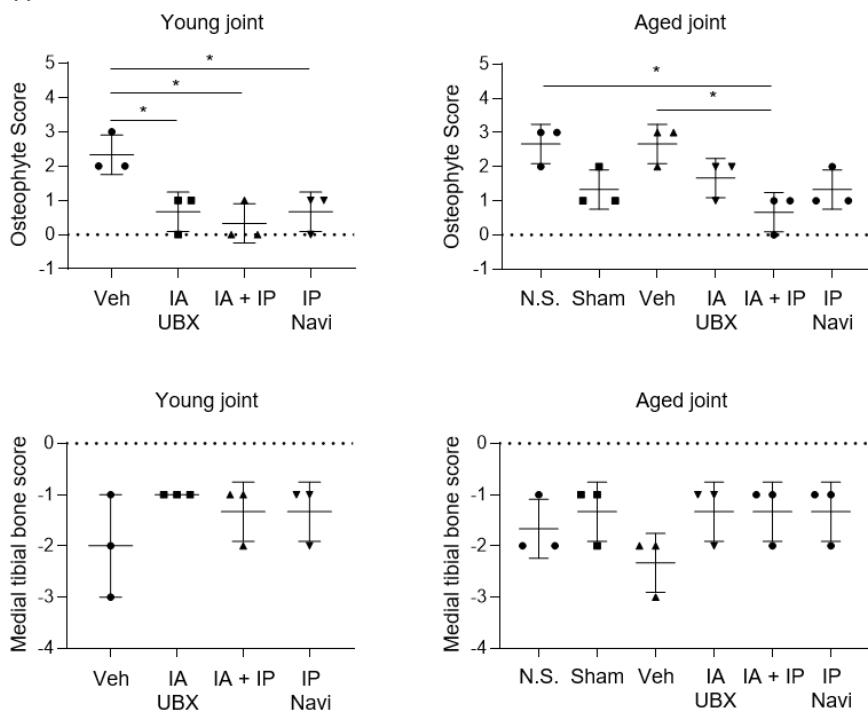
Supplemental Figure 9. IL-17 neutralization effects on joint gene expression. (A) Whole joint gene expression in young mice 4wks post ACLT with either isotype or IL-17 Nab administered (n=3). * $p<0.05$, and ** $p < 0.01$.



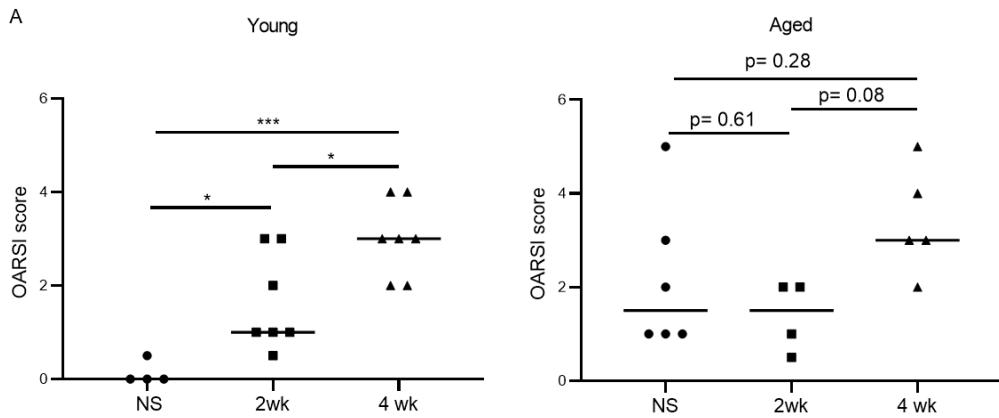
Supplemental Figure 10. Combined intra-articular treatment with navitoclax and UBX0101 does not enhance joint repair in aged mice. (A) Weight bearing of IA both group (IA injection of both UBX + Navi), (B) Representative images of Saf-O staining of IA both group joints, and OARSI score of IA both group joints. (C) CDKN2a gene expression of IA both group joints. Mice are age 76 weeks in all panels.

*** $p<0.001$. Experimental groups are compared to the control group (Veh).

A

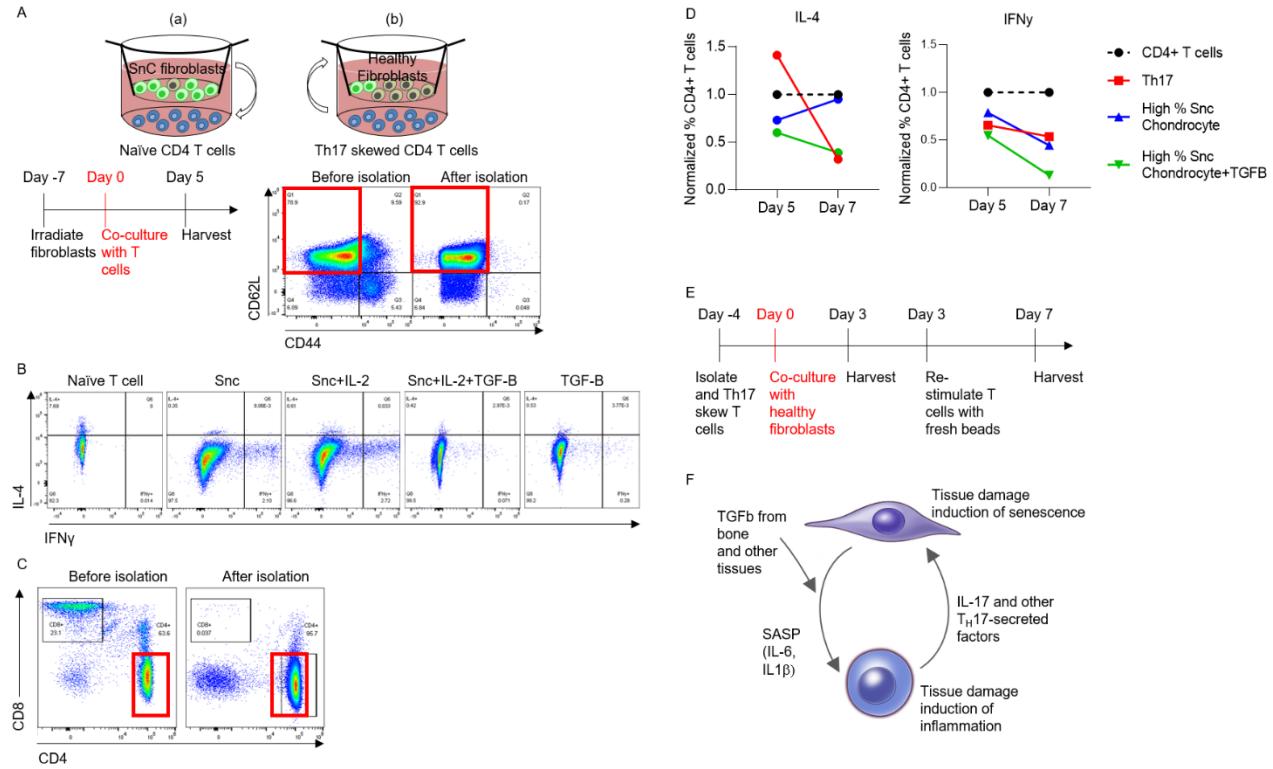


Supplemental Figure 11. (A) Osteophyte scoring on young and aged joints with IA UBX0101, IP navitoclax, or combined IA UBX0101 and IP navitoclax treatment after ACLT. * $p < 0.05$. Experimental groups are compared to the control group (Veh).

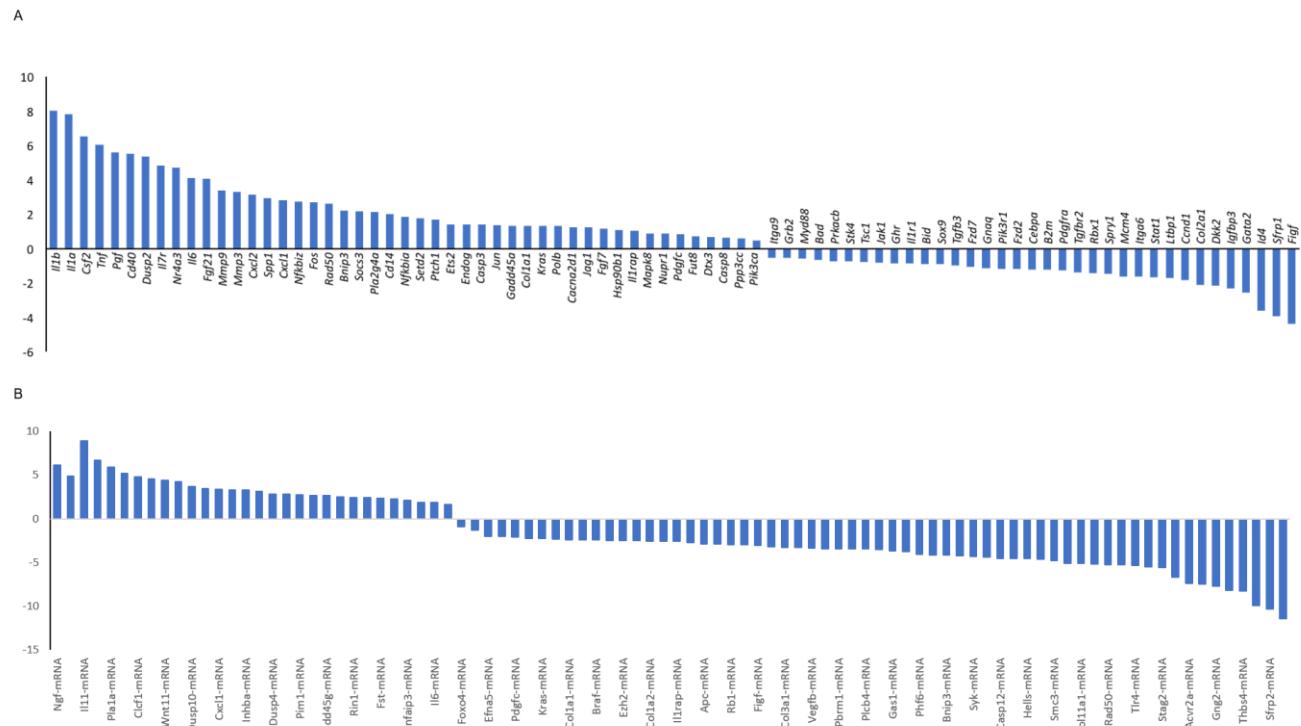


Supplemental Figure 12. ACLT induces joint damage as early as two weeks post injury. (A) Young or aged (76 weeks old) mice underwent ACL transection and were subsequently scored for cartilage damage using OARSI scoring at 2 and 4-weeks post ACLT. Mice 2wks post injury received no treatment or joint injections while mice 4wks post ACLT received one saline injection at 2wks post ACLT. * $p<0.05$.

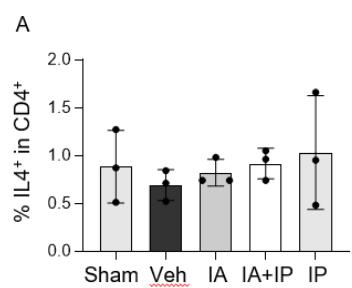
*** $p<0.001$.



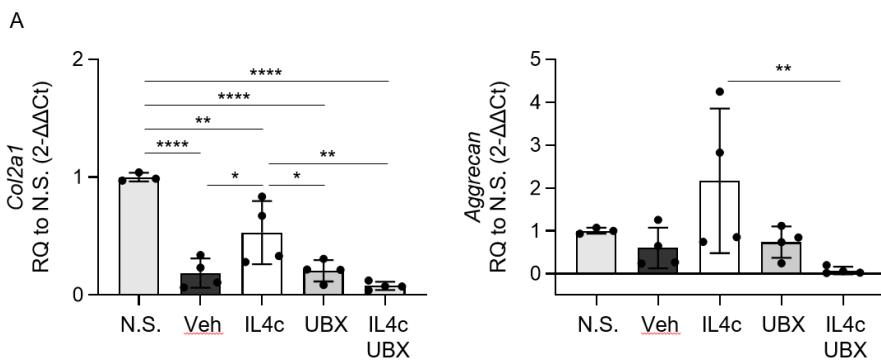
Supplemental Figure 13. Th17-senescent cell in vitro studies. (A) (a and b) Schematic of experimental set-up for in vitro experiments in figure 3 panels a and b. Experimental scheme and naïve CD4 T cell isolation from pooled lymph nodes before and after MACS isolation of naïve CD4 T cells. (B) Representative flow plots for IFNy quantification (C) Human CD4 T cell isolation from PMBCs. Left is PMBC, right is after MACS isolation of CD4 T cells. (D) Quantification of IL-4 and IFNy from human T cells after co-culture with Snc OA chondrocytes. Percentage of IL-4 or IFNy expressing CD4⁺ T cells were normalized to the control CD4⁺ T cell group within each time point. (E) Experimental scheme for Th17 cell co-culture with healthy fibroblasts. (F) Schematic of Th17 differentiation in the context of joint damage. Tissue damage induces senescent cell development, which supplies IL-6 and IL-1B needed for Th17 differentiation. Bone and other tissues provide TGF β , which is also needed for inducing Th17 differentiation. IL-17 and other factors secreted from Th17 cells then promote stromal cell senescence.



Supplemental Figure 14. Nanostring profiling of Th17 and irradiation induced senescence. (A) Complete list of significantly regulated genes in fibroblasts co-cultured with Th17 cells compared to normal fibroblasts. (B) Significantly regulated genes in irradiated fibroblasts compared to normal fibroblasts.



Supplemental Figure 15. Senolytic treatment does not enhance IL-4 protein levels. (A) Percentage of IL4+ cells in CD4+ T cells in inguinal lymph nodes in aged animals with senolytic treatment.



Supplemental Figure 16. Effect of IL-4 supplement in aged mice after ACL transection. (A) Whole joint gene expression 4 weeks after ACLT surgery. Surgery groups were treated with either Veh, IL-4 complex (IL4c), UBX or IL4c+UBX 2 weeks after ACLT surgery. * $p<0.05$, ** $p <0.01$, *** $p<0.001$ and **** $p<0.0001$.

S. Table 1 Mouse GO Th17 compared to healthy fibroblasts_downregulated

Enrichment Score:					
Annotation Cluster 1	Term	PValue	Genes	Fold Enrich	Bonferroni
Category					
UP_KEYWORDS	Signal	1.44E-08	FGFR2, TSPAN7, IGF1, GAS1, VEGFB, ACVR2A, ITGA6, SFRP1, SFRP2, COMP, NTRK2, EFNA5, NGFR, IGFBP3, LFNG, CSF1R	4.204039	9.38E-07
UP_KEYWORDS	Disulfide bond	2.73E-08	FGFR2, IGF1, VEGFB, ACVR2A, ITGA6, SFRP1, SFRP2, COMP, NTRK2, EFNA5, NGFR, IGFBP3, LFNG, CSF1R	5.349417	1.77E-06
UP_SEQ_FEATURE	disulfide bond	4.28E-07	FGFR2, VEGFB, ACVR2A, SFRP1, ITGA6, SFRP2, COMP, NTRK2, IGF1, EFNA5, NGFR, IGFBP3, CSF1R	4.90996	4.37E-05
UP_SEQ_FEATURE	signal peptide	4.51E-07	FGFR2, VEGFB, ACVR2A, SFRP1, ITGA6, SFRP2, COMP, NTRK2, IGF1, EFNA5, GAS1, NGFR, IGFBP3, CSF1R	4.248399	4.60E-05
Enrichment Score:					
Annotation Cluster 2	Term	PValue	Genes	Fold Enrich	Bonferroni
Category					
UP_KEYWORDS	Signal	1.44E-08	FGFR2, TSPAN7, IGF1, GAS1, VEGFB, ACVR2A, ITGA6, SFRP1, SFRP2, COMP, NTRK2, EFNA5, NGFR, IGFBP3, LFNG, CSF1R	4.204039	9.38E-07
UP_KEYWORDS	Glycoprotein glycosylation site:N-linked (GlcNAc...)	3.12E-07	FGFR2, TSPAN7, GAS1, VEGFB, ACVR2A, ITGA6, SFRP1, COMP, NTRK2, EFNA5, NGFR, IGFBP3, LFNG, CSF1R	4.380493	2.03E-05
UP_SEQ_FEATURE		1.97E-05	FGFR2, ACVR2A, SFRP1, ITGA6, COMP, NTRK2, TSPAN7, EFNA5, GAS1, NGFR, IGFBP3, LFNG, CSF1R	3.458883	0.002012
Enrichment Score:					
Annotation Cluster 3	Term	PValue	Genes	Fold Enrich	Bonferroni
Category	GO:0008284~positive regulation of cell proliferation	6.59E-06	FGFR2, SFRP1, SFRP2, NTRK2, IGF1, GAS1, CSF1R	12.97396	0.003551
GOTERM_BP_DIRECT					

	GO:0050680~negative regulation of epithelial cell proliferation	3.95E-05	FGFR2, SFRP1, SFRP2, GAS1	55.80864	0.02113
	GO:0050679~positive regulation of epithelial cell proliferation	4.65E-05	FGFR2, SFRP1, IGF1, GAS1	52.87135	0.024802
Enrichment Score:					
Annotation Cluster 4	4.3541011898918525				
Category Term PValue Genes Fold Enrich Bonferroni					
KEGG_PATHWAY	mmu04151:PI3K-Akt signaling pathway	2.94E-07	FGFR2, VEGFB, ITGA6, COMP, IGF1, EFNA5, NGFR, PIK3R1, CSF1R	10.95584	2.77E-05
KEGG_PATHWAY	mmu04015:Rap1 signaling pathway	4.14E-06	FGFR2, VEGFB, IGF1, EFNA5, NGFR, PIK3R1, CSF1R	13.97638	3.89E-04
KEGG_PATHWAY	mmu04014:Ras signaling pathway	6.13E-06	FGFR2, VEGFB, IGF1, EFNA5, NGFR, PIK3R1, CSF1R	13.06089	5.77E-04
GOTERM_BP_DIRECT	GO:0030335~positive regulation of cell migration	8.44E-04	ITGA6, IGF1, PIK3R1, CSF1R	19.7942	0.366194
KEGG_PATHWAY	mmu04510:Focal adhesion	9.21E-04	VEGFB, ITGA6, COMP, IGF1, PIK3R1	10.32072	0.082966
KEGG_PATHWAY	mmu05200:Pathways in cancer	0.00129	FGFR2, VEGFB, ITGA6, IGF1, PIK3R1, CSF1R	6.490295	0.114242
Enrichment Score:					
Annotation Cluster 5	3.4286934471703976				
Category Term PValue Genes Fold Enrich Bonferroni					
GOTERM_BP_DIRECT	GO:0043410~positive regulation of MAPK cascade	2.32E-06	FGFR2, NTRK2, IGF1, NGFR, IGFBP3	48.29594	0.001252
GOTERM_BP_DIRECT	GO:0007623~circadian rhythm	0.004534	NTRK2, IGF1, NGFR	27.90432	0.914028

GOTERM_BP_DIRECT	GO:0007399~nervous system development	0.00492 NTRK2, IGF1, EFNA5, NGFR	10.65841 0.930286
Enrichment Score: Annotation Cluster 6 3.340610858924582			
Category	Term	PValue Genes	Fold Enrich Bonferroni
UP_KEYWORDS	Developmental protein	7.43E-05 VEGFB, SFRP1, SFRP2, NTRK2, EFNA5, NGFR, LFNG	8.56126 0.004821
UP_KEYWORDS	Differentiation	1.16E-04 VEGFB, SFRP1, SFRP2, NTRK2, EFNA5, NGFR	11.08685 0.007525
GO:0007275~multicellular organism			
GOTERM_BP_DIRECT	development	2.41E-04 VEGFB, SFRP1, SFRP2, NTRK2, EFNA5, NGFR, LFNG	6.833711 0.121862
GOTERM_BP_DIRECT	GO:0030154~cell differentiation	5.92E-04 VEGFB, SFRP1, SFRP2, NTRK2, EFNA5, NGFR	7.72735 0.273492
UP_KEYWORDS	Neurogenesis	0.016112 NTRK2, EFNA5, NGFR	14.49819 0.6521
Enrichment Score: Annotation Cluster 7 2.471996764501841			
Category	Term	PValue Genes	Fold Enrich Bonferroni
GO:0008285~negative regulation of cell			
GOTERM_BP_DIRECT	proliferation	2.11E-05 FGFR2, SFRP1, SFRP2, IGF1, IGFBP3, CSF1R	15.69618 0.011328
GO:0050732~negative regulation of peptidyl-tyrosine			
GOTERM_BP_DIRECT	phosphorylation	1.12E-04 SFRP1, SFRP2, IGF1	177.2745 0.058798
GO:0045600~positive regulation of fat cell			
GOTERM_BP_DIRECT	differentiation	9.53E-04 SFRP1, SFRP2, IGF1	61.5034 0.40251
GO:0030307~positive regulation of cell growth			
GOTERM_BP_DIRECT	regulation of cell growth	0.003174 SFRP1, SFRP2, IGF1	33.48519 0.820323
GO:0005576~extracellular			
GOTERM_CC_DIRECT	ar region	0.003609 FGFR2, VEGFB, SFRP1, SFRP2, COMP, IGF1, IGFBP3	4.132285 0.253885

		GO:0001649~osteoblast differentiation	0.004952 SFRP1, IGF1, IGFBP3	26.66962	0.931468
UP_KEYWORDS	Secreted	GO:0005615~extracellular space	0.008507 VEGFB, SFRP1, SFRP2, COMP, IGF1, IGFBP3	4.250508	0.426112
GOTERM_CC_DIRECT	ar space	GO:0010629~negative regulation of gene expression	0.009599 VEGFB, SFRP1, SFRP2, COMP, IGF1, IGFBP3	4.128359	0.54216
GOTERM_BP_DIRECT	expression	GO:0070062~extracellular exosome	0.025175 SFRP1, SFRP2, IGF1	11.37233	0.999999
GOTERM_CC_DIRECT	ar exosome		0.724182 SFRP1, COMP, IGFBP3	1.161005	1
Enrichment Score:					
Annotation Cluster 8	2.208186783569238				
Category	Term	PValue	Genes	Fold Enrich Bonferroni	
KEGG_PATHWAY	mmu05200:Pathways in cancer	0.00129	FGFR2, VEGFB, ITGA6, IGF1, PIK3R1, CSF1R	6.490295	0.114242
GO:0045944~positive regulation of transcription from RNA polymerase II promoter					
GOTERM_BP_DIRECT	mmu04550:Signaling pathways regulating pluripotency of stem cells	0.001773	FGFR2, ACVR2A, ITGA6, SFRP2, IGF1, PIK3R1	6.057621	0.616461
KEGG_PATHWAY	mmu05215:Prostate cancer	0.003197	FGFR2, ACVR2A, IGF1, PIK3R1	12.38486	0.259891
KEGG_PATHWAY	mmu04810:Regulation of actin cytoskeleton	0.015746	FGFR2, IGF1, PIK3R1	14.56629	0.775052
KEGG_PATHWAY		0.079067	FGFR2, ITGA6, PIK3R1	6.017997	0.999566
Enrichment Score:					
Annotation Cluster 9	2.063396684684468				
Category	Term	PValue	Genes	Fold Enrich Bonferroni	

	GO:0006468~protein phosphorylation	9.33E-06 FGFR2, ACVR2A, MAP3K1, NTRK2, IGFBP3, PIK3R1, CSF1R	12.20814	0.005028
GOTERM_BP_DIRECT	GO:0009986~cell			
GOTERM_CC_DIRECT	surface	1.40E-05 FGFR2, ACVR2A, SFRP1, ITGA6, NTRK2, NGFR, CSF1R	11.51653	0.001132
UP_KEYWORDS	Kinase	1.77E-04 FGFR2, ACVR2A, MAP3K1, NTRK2, PIK3R1, CSF1R	10.13028	0.011464
	GO:0016301~kinase			
GOTERM_MF_DIRECT	activity	3.56E-04 FGFR2, ACVR2A, MAP3K1, NTRK2, PIK3R1, CSF1R	8.628091	0.037405
	IPR000719:Protein kinase, catalytic domain	8.95E-04 FGFR2, ACVR2A, MAP3K1, NTRK2, CSF1R	10.52325	0.070783
INTERPRO	IPR011009:Protein kinase-like domain	0.00119 FGFR2, ACVR2A, MAP3K1, NTRK2, CSF1R	9.747255	0.093001
	GO:0004714~transmembrane receptor protein tyrosine kinase activity	0.001241 FGFR2, NTRK2, CSF1R	53.84568	0.124455
UP_KEYWORDS	Transferase	0.001288 FGFR2, ACVR2A, MAP3K1, NTRK2, LFNG, PIK3R1, CSF1R	5.051868	0.080343
UP_SEQ_FEATURE	domain:Protein kinase	0.001337 FGFR2, ACVR2A, MAP3K1, NTRK2, CSF1R	9.442231	0.127534
	GO:0004672~protein kinase activity	0.001473 FGFR2, ACVR2A, MAP3K1, NTRK2, CSF1R	9.126386	0.145886
GOTERM_MF_DIRECT	GO:0016310~phosphorylation	0.002175 FGFR2, ACVR2A, MAP3K1, NTRK2, CSF1R	8.207153	0.691497
GOTERM_BP_DIRECT	IPR020635:Tyrosine-protein kinase, catalytic domain	0.002244 FGFR2, NTRK2, CSF1R	40.14425	0.168252
UP_SEQ_FEATURE	binding site:ATP	0.002315 FGFR2, ACVR2A, MAP3K1, NTRK2, CSF1R	8.13036	0.210517
	IPR008266:Tyrosine-protein kinase, active site	0.003329 FGFR2, NTRK2, CSF1R	32.8453	0.239228
INTERPRO	Tyrosine-protein kinase	0.003573 FGFR2, NTRK2, CSF1R	31.69073	0.207595
UP_KEYWORDS				

	IPR017441:Protein kinase, ATP binding site active site:Proton	0.00458 FGFR2, MAP3K1, NTRK2, CSF1R	11.00401	0.313697
INTERPRO				
UP_SEQ_FEATURE	acceptor	0.004709 FGFR2, ACVR2A, MAP3K1, NTRK2, CSF1R	6.676056	0.382124
SMART	SM00219:TyrKc	0.005863 FGFR2, NTRK2, CSF1R	24.13194	0.121355
	GO:0004713~protein tyrosine kinase activity	0.006061 FGFR2, NTRK2, CSF1R	24.0303	0.478224
GOTERM_MF_DIRECT	GO:0042803~protein homodimerization			
GOTERM_MF_DIRECT	activity	0.006416 FGFR2, VEGFB, NTRK2, NGFR, CSF1R	6.072821	0.497776
	IPR001245:Serine-threonine/tyrosine-protein kinase catalytic domain	0.006447 FGFR2, NTRK2, CSF1R	23.39341	0.41161
UP_SEQ_FEATURE	domain:Ig-like C2-type 1	0.007552 FGFR2, NTRK2, CSF1R	21.54545	0.538478
UP_SEQ_FEATURE	domain:Ig-like C2-type 2	0.007663 FGFR2, NTRK2, CSF1R	21.38346	0.543703
GOTERM_MF_DIRECT	GO:0016740~transferas e activity	0.011116 FGFR2, ACVR2A, MAP3K1, NTRK2, LFNG, CSF1R	3.950634	0.697631
GOTERM_BP_DIRECT	GO:0046777~protein autophosphorylation	0.012538 FGFR2, NTRK2, CSF1R	16.46812	0.998901
GOTERM_BP_DIRECT	GO:0001934~positive regulation of protein phosphorylation	0.012668 ACVR2A, PIK3R1, CSF1R	16.37862	0.998977
UP_SEQ_FEATURE	nucleotide phosphate-binding region:ATP	0.013615 FGFR2, ACVR2A, MAP3K1, NTRK2, CSF1R	4.922118	0.752968
INTERPRO	IPR003598:Immunoglob ulin subtype 2	0.018589 FGFR2, NTRK2, CSF1R	13.43671	0.785322
UP_SEQ_FEATURE	topological domain:Extracellular	0.018684 FGFR2, ACVR2A, ITGA6, NTRK2, TSPAN7, NGFR, CSF1R	2.941489	0.853949
UP_KEYWORDS	ATP-binding	0.020178 FGFR2, ACVR2A, MAP3K1, NTRK2, CSF1R	4.378886	0.7342

UP_KEYWORDS	Nucleotide-binding	0.045484 FGFR2, ACVR2A, MAP3K1, NTRK2, CSF1R	3.402749	0.951482
SMART	SM00408:IGc2	0.046193 FGFR2, NTRK2, CSF1R	8.077221	0.64671
UP_KEYWORDS	Receptor GO:0010628~positive regulation of gene expression	0.048042 FGFR2, ACVR2A, ITGA6, NTRK2, NGFR, CSF1R	2.740951	0.959247
GOTERM_BP_DIRECT	GO:0005524~ATP binding	0.053101 FGFR2, NTRK2, PIK3R1	7.553049	1
GOTERM_MF_DIRECT	binding	0.053144 FGFR2, ACVR2A, MAP3K1, NTRK2, CSF1R	3.215734	0.9971
UP_KEYWORDS	Immunoglobulin domain	0.054902 FGFR2, NTRK2, CSF1R	7.445016	0.974533
INTERPRO	IPR003599:Immunoglobulin subtype	0.074108 FGFR2, NTRK2, CSF1R	6.277383	0.998189
KEGG_PATHWAY	mmu04010:MAPK signaling pathway	0.104663 FGFR2, MAP3K1, NTRK2	5.106906	0.999969
GOTERM_MF_DIRECT	GO:0000166~nucleotide binding	0.111309 FGFR2, ACVR2A, MAP3K1, NTRK2, CSF1R	2.503157	0.999997
UP_KEYWORDS	Ubl conjugation	0.113048 FGFR2, NTRK2, PIK3R1, CSF1R	3.172583	0.999589
UP_KEYWORDS	Cell membrane	0.165319 FGFR2, ITGA6, NTRK2, EFNA5, GAS1, CSF1R	1.905322	0.999992
SMART	SM00409:IG	0.169195 FGFR2, NTRK2, CSF1R	3.773528	0.983057
INTERPRO	IPR007110:Immunoglobulin-like domain	0.190942 FGFR2, NTRK2, CSF1R	3.534439	1
INTERPRO	IPR013783:Immunoglobulin-like fold	0.249226 FGFR2, NTRK2, CSF1R	2.958766	1
GOTERM_CC_DIRECT	GO:0005737~cytoplasm	0.601574 FGFR2, ACVR2A, MAP3K1, NTRK2, IGF1, NGFR, PIK3R1	1.092429	1
Enrichment Score:				
Annotation Cluster 10	1.8193059244053358			
Category	Term	PValue	Genes	Fold Enrich Bonferroni
	GO:0009986~cell			
GOTERM_CC_DIRECT	surface	1.40E-05	FGFR2, ACVR2A, SFRP1, ITGA6, NTRK2, NGFR, CSF1R	11.51653 0.001132
	glycosylation site:N-linked (GlcNAc...)		FGFR2, ACVR2A, SFRP1, ITGA6, COMP, NTRK2, TSPAN7, EFNA5, GAS1, NGFR, IGFBP3, LFNG, CSF1R	
UP_SEQ_FEATURE		1.97E-05		3.458883 0.002012

	GO:0005886~plasma membrane	FGFR2, ACVR2A, SFRP1, ITGA6, NTRK2, IGF1, EFNA5, GAS1, NGFR, PIK3R1, CSF1R			
GOTERM_CC_DIRECT	GO:0016020~membrane	0.005059 FGFR2, VEGFB, ACVR2A, ITGA6, MAP3K1, NTRK2, TSPAN7, EFNA5, GAS1, NGFR, LFNG, PIK3R1, CSF1R	2.335507	0.336899	
GOTERM_CC_DIRECT	e topological	0.007171 FGFR2, ACVR2A, ITGA6, NTRK2, TSPAN7, NGFR, LFNG, PIK3R1, CSF1R	1.922399	0.441724	
UP_SEQ_FEATURE	domain:Cytoplasmic topological	0.016563 FGFR2, ACVR2A, ITGA6, NTRK2, TSPAN7, NGFR, LFNG, CSF1R	2.633333	0.817966	
UP_SEQ_FEATURE	domain:Extracellular	0.018684 FGFR2, ACVR2A, ITGA6, NTRK2, TSPAN7, NGFR, CSF1R	2.941489	0.853949	
UP_KEYWORDS	Receptor GO:0016021~integral component of membrane	0.048042 FGFR2, ACVR2A, ITGA6, NTRK2, NGFR, CSF1R FGFR2, ACVR2A, SFRP1, ITGA6, SFRP2, NTRK2, TSPAN7, NGFR, LFNG, PIK3R1, CSF1R	2.740951	0.959247	
GOTERM_CC_DIRECT	GO:0005887~integral component of plasma membrane	0.059403 NGFR, LFNG, PIK3R1, CSF1R	1.655025	0.99299	
GOTERM_CC_DIRECT	membrane	0.080416 FGFR2, NTRK2, TSPAN7, NGFR FGFR2, ACVR2A, ITGA6, NTRK2, TSPAN7, EFNA5, GAS1,	3.676171	0.998876	
UP_KEYWORDS	Membrane	0.10416 NGFR, LFNG, PIK3R1, CSF1R	1.512211	0.999215	
UP_SEQ_FEATURE	transmembrane region	0.115732 FGFR2, ACVR2A, ITGA6, NTRK2, TSPAN7, NGFR, LFNG, CSF1R FGFR2, ACVR2A, ITGA6, NTRK2, TSPAN7, NGFR, LFNG,	1.758813	0.999996	
UP_KEYWORDS	Transmembrane helix	0.15371 PIK3R1, CSF1R FGFR2, ACVR2A, ITGA6, NTRK2, TSPAN7, NGFR, LFNG,	1.548452	0.999981	
UP_KEYWORDS	Transmembrane	0.155423 PIK3R1, CSF1R	1.544667	0.999983	
UP_KEYWORDS	Cell membrane	0.165319 FGFR2, ITGA6, NTRK2, EFNA5, GAS1, CSF1R	1.905322	0.999992	

Enrichment Score:

Annotation Cluster 11 1.110395136017742

Category	Term	PValue	Genes	Fold Enrich	Bonferroni
	mmu05200:Pathways in				
KEGG_PATHWAY	cancer	0.00129	FGFR2, VEGFB, ITGA6, IGF1, PIK3R1, CSF1R	6.490295	0.114242
UP_KEYWORDS	Cell membrane	0.165319	FGFR2, ITGA6, NTRK2, EFNA5, GAS1, CSF1R	1.905322	0.999992
UP_KEYWORDS	Alternative splicing	0.324713	FGFR2, VEGFB, ITGA6, NTRK2, IGF1, EFNA5	1.498662	1
UP_SEQ_FEATURE	splice variant	0.522521	FGFR2, VEGFB, ITGA6, NTRK2, IGF1, EFNA5	1.212535	1

S. Table 1 Mouse GO Th17 compared to healthy fibroblasts_upregulated

Enrichment Score:					
Annotation Cluster 1	Term	PValue	Genes	Fold Enrich	Bonferroni
Category	Growth factor	2.34E-09	CXCL1, LIF, INHBA, IL6, PGF, FGF2, IL11, NGF	34.8454	2.34E-07
UP_KEYWORDS	GO:0008083~growth factor				
GOTERM_MF_DIRECT	activity	3.51E-08	CXCL1, LIF, INHBA, IL6, PGF, FGF2, IL11, NGF	23.47653	5.23E-06
GOTERM_CC_DIRECT	GO:0005615~extracellular space	5.38E-07	CXCL1, IL6, WNT5B, PGF, TNC, MMP9, MMP3, IL11, LIF, INHBA, CLCF1, HSPB1, WNT11, FGF2, NGF	4.902427	4.41E-05
UP_KEYWORDS	Secreted	5.44E-07	CXCL1, IL6, WNT5B, PGF, TNC, MMP9, MMP3, IL11, LIF, INHBA, CLCF1, PLA1A, WNT11, FGF2, NGF	4.924369	5.44E-05
GOTERM_CC_DIRECT	GO:0005576~extracellular region	3.38E-06	CXCL1, IL6, WNT5B, PGF, TNC, MMP9, MMP3, IL11, LIF, INHBA, CLCF1, PLA1A, WNT11, FGF2, NGF	4.206075	2.77E-04
LIF, DVL3, INHBA, IL6, WNT5B, CLCF1, WNT11,					
GOTERM_MF_DIRECT	GO:0005102~receptor binding	3.58E-06	FGF2, NGF	9.295169	5.34E-04
GOTERM_MF_DIRECT	GO:0005125~cytokine activity	8.61E-06	CXCL1, LIF, INHBA, IL6, CLCF1, FGF2, IL11	13.91862	0.001282
CXCL1, CACNA2D1, IL6, WNT5B, PGF, TNC, MMP9, MMP3, IL11, LIF, INHBA, CLCF1, CSF2RB, PLA1A,					
UP_SEQ_FEATURE	signal peptide	0.001423	WNT11, NGF	2.306274	0.246792
UP_KEYWORDS	Disulfide bond	0.001989	CXCL1, CACNA2D1, IL6, WNT5B, PGF, MMP9, TNC, MMP3, LIF, INHBA, CSF2RB, PLA1A, WNT11, NGF	2.478998	0.180568
CXCL1, CACNA2D1, IL6, WNT5B, PGF, TNC, MMP9, MMP3, IL11, LIF, INHBA, CLCF1, CSF2RB, PLA1A,					
UP_KEYWORDS	Signal	0.00798	WNT11, NGF	1.948213	0.551225
CACNA2D1, WNT5B, PGF, MMP9, TNC, MMP3, LIF, INHBA, CLCF1, CSF2RB, CREB3L1, PLA1A, WNT11,					
UP_KEYWORDS	Glycoprotein	0.011342	NGF	2.029984	0.680395
CACNA2D1, WNT5B, PGF, MMP9, TNC, MMP3, LIF, INHBA, CLCF1, CSF2RB, CREB3L1, PLA1A, WNT11,					
UP_SEQ_FEATURE	glycosylation site:N-linked (GlcNAc...)	0.032568	NGF	1.769352	0.998625

UP_SEQ_FEATURE	disulfide bond	CXCL1, LIF, INHBA, IL6, PGF, MMP9, TNC, CSF2RB, 0.03747 PLA1A, MMP3, NGF	1.973426	0.999499
Enrichment Score:				
Annotation Cluster 2 3.297593221134542				
Category	Term	PValue	Genes	Fold Enrich Bonferroni
KEGG_PATHWAY	mmu04668:TNF signaling pathway	3.13E-09	CXCL1, LIF, FOS, IL6, SOCS3, MMP9, NFKBIA, CREB3L1, TNFAIP3, MMP3	17.20967 4.14E-07
KEGG_PATHWAY	mmu05164:Influenza A	0.011562	PRKCA, IL6, HSPA2, SOCS3, NFKBIA	5.484952 0.784574
KEGG_PATHWAY	mmu04931:Insulin resistance	0.019165	IL6, SOCS3, NFKBIA, CREB3L1	6.821286 0.922255
KEGG_PATHWAY	mmu05168:Herpes simplex infection	0.092929	FOS, IL6, SOCS3, NFKBIA	3.607411 0.999997
Enrichment Score:				
Annotation Cluster 3 2.7755941169736236				
Category	Term	PValue	Genes	Fold Enrich Bonferroni
KEGG_PATHWAY	mmu04916:Melanogenesis	1.47E-04	PRKCA, DVL3, WNT5B, LEF1, CREB3L1, WNT11	11.36881 0.019218
GOTERM_BP_DIRECT	GO:0001649~osteoblast differentiation	0.001984	TNC, LEF1, CREB3L1, WNT11	15.61148 0.769549
GOTERM_MF_DIRECT	GO:0044212~transcription regulatory region DNA binding	0.016156	FOS, LEF1, CREB3L1, WNT11	7.30493 0.911695
Enrichment Score:				
Annotation Cluster 4 2.636553309813029				
Category	Term	PValue	Genes	Fold Enrich Bonferroni
GOTERM_MF_DIRECT	GO:0005125~cytokine activity	8.61E-06	CXCL1, LIF, INHBA, IL6, CLCF1, FGF2, IL11	13.91862 0.001282
UP_KEYWORDS	Cytokine	4.18E-04	CXCL1, LIF, IL6, CLCF1, IL11	13.82927 0.040973
GOTERM_BP_DIRECT	GO:0046888~negative regulation of hormone secretion	4.27E-04	LIF, IL6, IL11	94.50523 0.270646
INTERPRO	IPR009079:Four-helical cytokine-like, core	7.33E-04	LIF, IL6, CLCF1, IL11	22.0788 0.09486

KEGG_PATHWAY	mmu04630:Jak-STAT signaling pathway	8.60E-04 LIF, IL6, SOCS3, PIM1, CSF2RB, IL11	7.762153	0.107321
KEGG_PATHWAY	mmu04060:Cytokine-cytokine receptor interaction	0.001426 CXCL1, LIF, INHBA, IL6, CLCF1, CSF2RB, IL11	5.426023	0.171709
GOTERM_BP_DIRECT	GO:0042517~positive regulation of tyrosine phosphorylation of Stat3	0.001996 LIF, IL6, CLCF1	44.10244	0.771541
GOTERM_BP_DIRECT	GO:0033138~positive regulation of peptidyl-serine phosphorylation	0.011656 LIF, IL6, IL11	17.87937	0.999827
INTERPRO	IPR012351:Four-helical cytokine, core	0.012673 LIF, IL6, CLCF1	17.12361	0.823528
GOTERM_BP_DIRECT	GO:0050731~positive regulation of peptidyl-tyrosine phosphorylation	0.020993 LIF, IL6, IL11	13.09973	1
GOTERM_BP_DIRECT	GO:0043410~positive regulation of MAPK cascade	0.022172 LIF, IL6, IL11	12.72186	1
GOTERM_BP_DIRECT	GO:0006955~immune response	0.121302 CXCL1, LIF, IL6	4.86424	1

Enrichment Score:

Annotation Cluster 5 2.4746683250657333

Category	Term	PValue	Genes	Fold Enrich	Bonferroni
GOTERM_BP_DIRECT	GO:0045766~positive regulation of angiogenesis	1.45E-04	PRKCA, PGF, MMP9, HSPB1, FGF2	18.22415	0.101437
KEGG_PATHWAY	mmu04151:PI3K-Akt signaling pathway	3.70E-04	PRKCA, IL6, PGF, TNC, NR4A1, CREB3L1, FGF2, DDIT4, NGF	4.809881	0.047696
GOTERM_BP_DIRECT	GO:0001938~positive regulation of endothelial cell proliferation	4.36E-04	PRKCA, PGF, NR4A1, FGF2	26.32981	0.275242
KEGG_PATHWAY	mmu04014:Ras signaling pathway	0.001072	PRKCA, REL, PGF, RIN1, PLA1A, FGF2, NGF	5.73405	0.132048
UP_KEYWORDS	Angiogenesis	0.019593	PRKCA, PGF, FGF2	13.60256	0.861757

GOTERM_BP_DIRECT	GO:0001525~angiogenesis mmu04015:Rap1 signaling pathway	0.097874 PRKCA, PGF, FGF2	5.535871	1
KEGG_PATHWAY		0.099149 PRKCA, PGF, FGF2, NGF	3.506269	0.999999
Enrichment Score:				
Annotation Cluster 6 2.388996644745089				
Category	Term	PValue Genes	Fold Enrich	Bonferroni
UP_KEYWORDS	Extracellular matrix GO:0030335~positive	7.66E-04 WNT5B, MMP9, TNC, WNT11, MMP3	11.76959	0.073784
GOTERM_BP_DIRECT	regulation of cell migration GO:0005578~proteinaceous	0.001027 PRKCA, WNT5B, LEF1, WNT11, MMP3	10.86267	0.53209
GOTERM_CC_DIRECT	extracellular matrix GO:0031012~extracellular	0.003457 WNT5B, MMP9, TNC, WNT11, MMP3	7.77769	0.247206
GOTERM_CC_DIRECT	matrix mmu05205:Proteoglycans in	0.020341 MMP9, TNC, HSPB1, MMP3	6.687755	0.814592
KEGG_PATHWAY	cancer	0.020509 PRKCA, WNT5B, MMP9, WNT11, FGF2	4.620329	0.935132
Enrichment Score:				
Annotation Cluster 7 2.384409938694051				
Category	Term	PValue Genes	Fold Enrich	Bonferroni
KEGG_PATHWAY	mmu04310:Wnt signaling pathway	PRKCA, DVL3, WNT5B, LEF1, WNT11, FOSL1, 7.84E-05 NFATC1	9.312749	0.010297
KEGG_PATHWAY	mmu04916:Melanogenesis	1.47E-04 PRKCA, DVL3, WNT5B, LEF1, CREB3L1, WNT11	11.36881	0.019218
KEGG_PATHWAY	mmu04550:Signaling pathways regulating pluripotency of stem cells GO:0030335~positive	6.87E-04 LIF, DVL3, INHBA, WNT5B, WNT11, FGF2	8.155885	0.08671
GOTERM_BP_DIRECT	regulation of cell migration mmu05217:Basal cell carcinoma	0.001027 PRKCA, WNT5B, LEF1, WNT11, MMP3	10.86267	0.53209
GOTERM_MF_DIRECT	GO:0005109~frizzled binding	0.002691 DVL3, WNT5B, LEF1, WNT11	13.89521	0.299271
UP_KEYWORDS	Wnt signaling pathway GO:0016055~Wnt signaling pathway	0.00342 DVL3, WNT5B, WNT11	33.59307	0.399791
GOTERM_BP_DIRECT		0.003676 DVL3, WNT5B, LEF1, WNT11	12.57206	0.308072
GOTERM_BP_DIRECT		0.011558 DVL3, WNT5B, LEF1, WNT11	8.282148	0.999814

GOTERM_BP_DIRECT	GO:0060070~canonical Wnt signaling pathway	0.016551 DVL3, LEF1, WNT11	14.86599	0.999996	
GOTERM_BP_DIRECT	GO:0090090~negative regulation of canonical Wnt signaling pathway	0.021383 WNT5B, LEF1, WNT11	12.97131	1	
UP_KEYWORDS	Developmental protein mmu04390:Hippo signaling pathway	0.027624 DVL3, WNT5B, PGF, CREB3L1, WNT11, FGF2	3.40064	0.939264	
KEGG_PATHWAY	GO:0007275~multicellular organism development	0.043193 DVL3, WNT5B, LEF1, WNT11	4.969149	0.997057	
GOTERM_BP_DIRECT	Enrichment Score: 2.3427938579208942	0.074936 DVL3, WNT5B, PGF, CREB3L1, WNT11, FGF2	2.571571	1	
Annotation Cluster 8	Category	Term	PValue	Genes	Fold Enrich Bonferroni
		GO:0001077~transcriptional activator activity, RNA polymerase II core promoter proximal region sequence-specific binding		FOS, LEF1, NR4A1, CREB3L1, HMGA2, FOSL1,	
GOTERM_MF_DIRECT		IPR004827:Basic-leucine zipper domain	3.21E-05	NFATC1	11.0318 0.004767
INTERPRO		SMART	0.004938	FOS, CREB3L1, FOSL1	27.90515 0.489971
UP_SEQ_FEATURE		IPR008343:Mitogen-activated protein (MAP) kinase phosphatase	0.011912	FOS, CREB3L1, FOSL1	17.52101 0.380811
UP_SEQ_FEATURE		domain:Leucine-zipper	0.022849	FOS, CREB3L1, FOSL1	12.50833 0.989946
UP_SEQ_FEATURE		DNA-binding region:Basic motif	0.044818	FOS, CREB3L1, FOSL1	8.659615 0.999891
Annotation Cluster 9	Category	Term	PValue	Genes	Fold Enrich Bonferroni
		IPR008343:Mitogen-activated protein (MAP) kinase phosphatase	1.64E-04	DUSP5, DUSP4, DUSP10	150.6878 0.022047

	GO:0017017~MAP kinase tyrosine-serine/threonine phosphatase activity	3.93E-04 DUSP5, DUSP4, DUSP10	98.19512	0.056953
GOTERM_MF_DIRECT	IPR001763:Rhodanese-like domain	9.88E-04 DUSP5, DUSP4, DUSP10	62.78659	0.125809
INTERPRO	IPR024950:Dual specificity phosphatase	0.001651 DUSP5, DUSP4, DUSP10	48.60897	0.201217
INTERPRO	IPR020422:Dual specificity phosphatase, subgroup, catalytic domain	0.002102 DUSP5, DUSP4, DUSP10	43.05366	0.248837
SMART	SM00450:RHOD	0.002502 DUSP5, DUSP4, DUSP10	38.85093	0.095362
INTERPRO	IPR000340:Dual specificity phosphatase, catalytic domain	0.002738 DUSP5, DUSP4, DUSP10	37.67195	0.311283
	GO:0008138~protein tyrosine/serine/threonine phosphatase activity			
GOTERM_MF_DIRECT	IPR016130:Protein-tyrosine phosphatase, active site	0.003784 DUSP5, DUSP4, DUSP10	31.91341	0.431559
SMART	SM00195:DSPc	0.005743 DUSP5, DUSP4, DUSP10	25.53061	0.205763
INTERPRO	IPR000387:Protein-tyrosine/Dual specificity phosphatase	0.008861 DUSP5, DUSP4, DUSP10	20.64217	0.701948
INTERPRO	GO:0004725~protein tyrosine phosphatase activity	0.011076 DUSP5, DUSP4, DUSP10	18.37656	0.780121
GOTERM_MF_DIRECT	UP_KEYWORDS	0.020807 DUSP5, DUSP4, DUSP10	13.16017	0.956411
	Protein phosphatase	0.022063 DUSP5, DUSP4, DUSP10	12.76548	0.892575
GOTERM_BP_DIRECT	GO:0016311~dephosphorylation	0.024199 DUSP5, DUSP4, DUSP10	12.13829	1
GOTERM_MF_DIRECT	on	0.029935 DUSP5, DUSP4, DUSP10	10.81811	0.989202
GOTERM_MF_DIRECT	GO:0016791~phosphatase activity	0.040364 DUSP5, DUSP4, DUSP10	9.183716	0.997843
GOTERM_MF_DIRECT	GO:0004721~phosphoprotein phosphorylation	DUSP5, DUSP4, MMP9, DUSP10, PLA1A, TNFAIP3,		
UP_KEYWORDS	Hydrolase	0.066808 MMP3	2.352488	0.999007

Enrichment Score:					
Annotation Cluster	Term	PValue	Genes	Fold Enrich	Bonferroni
Annotation Cluster 10	2.246166125995958				
Category	GO:0045944~positive regulation of transcription from RNA polymerase II promoter	1.29E-08	IL6, NFKBIA, NR4A1, LEF1, HMGA2, IL11, LIF, INHBA, FOS, REL, CREB3L1, FOSL1, FGF2, ETV4, NFATC1	6.648609	9.53E-06
GOTERM_BP_DIRECT	GO:0001077~transcriptional activator activity, RNA polymerase II core promoter proximal region sequence-specific binding	3.21E-05	FOS, LEF1, NR4A1, CREB3L1, HMGA2, FOSL1, NFATC1	11.0318	0.004767
GOTERM_MF_DIRECT	Nucleus	2.99E-04	PRKCA, NFKBIZ, PIM1, DUSP10, NFKBIA, NR4A1, LEF1, LIG4, HMGA2, FOS, DUSP4, REL, CREB3L1, HSPB1, TNFAIP3, FOSL1, FGF2, ETV4, NFATC1	2.318095	0.029441
UP_KEYWORDS	GO:0043565~sequence-specific DNA binding	5.26E-04	FOS, REL, LEF1, NR4A1, CREB3L1, FOSL1, ETV4, NFATC1	5.377721	0.07538
GOTERM_MF_DIRECT	DNA-binding	0.001596	FOSL1, ETV4, NFATC1	3.448695	0.147598
UP_KEYWORDS	GO:0006357~regulation of transcription from RNA polymerase II promoter	0.001739	FOS, NFKBIZ, INHBA, LEF1, FOSL1, NFATC1	6.665356	0.723776
GOTERM_BP_DIRECT	GO:0005667~transcription factor complex	0.001882	FOS, REL, LEF1, NR4A1, NFATC1	9.205056	0.143129
GOTERM_CC_DIRECT	GO:0003700~transcription factor activity, sequence-specific DNA binding	0.003583	FOS, REL, LEF1, NR4A1, CREB3L1, FOSL1, ETV4, NFATC1	3.85515	0.414198
GOTERM_MF_DIRECT	GO:0005634~nucleus	0.005544	PRKCA, NFKBIZ, PIM1, DUSP10, NFKBIA, LEF1, NR4A1, LIG4, HMGA2, MMP3, FOS, DUSP4, REL, HSPA2, HSPB1, CREB3L1, TNFAIP3, FOSL1, FGF2, ETV4, NFATC1	1.714994	0.366124
GOTERM_CC_DIRECT	GO:0003677~DNA binding	0.007482	FOS, REL, LEF1, NR4A1, CREB3L1, LIG4, HMGA2, TNFAIP3, FOSL1, ETV4, NFATC1	2.534182	0.673416

	GO:0008134~transcription factor binding	0.007606 FOS, PIM1, LEF1, NR4A1, NFATC1	6.220939	0.679444
GOTERM_MF_DIRECT	GO:0000978~RNA polymerase II core promoter proximal region sequence-specific DNA binding	0.00899 FOS, LEF1, FOSL1, ETV4, NFATC1	5.926354	0.73959
GOTERM_MF_DIRECT	GO:0000977~RNA polymerase II regulatory region sequence-specific DNA binding	0.014038 REL, LEF1, FOSL1, NFATC1	7.701578	0.878344
GOTERM_MF_DIRECT	GO:0003682~chromatin binding	0.021513 FOS, REL, LEF1, CREB3L1, NFATC1	4.565581	0.960853
UP_KEYWORDS	Activator	0.023715 NFKBIZ, LEF1, CREB3L1, ETV4, NFATC1 NFKBIZ, REL, LEF1, NR4A1, CREB3L1, HMGA2, ETV4,	4.432458	0.909288
UP_KEYWORDS	Transcription regulation	0.036043 NFATC1 NFKBIZ, REL, LEF1, NR4A1, CREB3L1, HMGA2, ETV4,	2.459903	0.974545
UP_KEYWORDS	Transcription	0.041957 NFATC1 FOS, NFKBIZ, REL, LEF1, NR4A1, CREB3L1, HMGA2,	2.380509	0.986244
GOTERM_BP_DIRECT	GO:0006355~regulation of transcription, DNA-templated	0.057428 FOSL1, ETV4, NFATC1	1.935166	1
GOTERM_BP_DIRECT	GO:0006351~transcription, DNA-templated	0.117251 NFATC1 NFKBIZ, REL, LEF1, NR4A1, CREB3L1, HMGA2, ETV4,	1.871722	1
GOTERM_BP_DIRECT	GO:0000122~negative regulation of transcription from RNA polymerase II promoter	0.217659 REL, LEF1, HMGA2, NFATC1 NFKBIZ, CACNA2D1, TNC, LEF1, CREB3L1, HSPB1,	2.419887	1
UP_KEYWORDS	Alternative splicing	0.767575 ETV4, NFATC1 NFKBIZ, CACNA2D1, TNC, PIM1, LEF1, CREB3L1,	0.926002	1
UP_SEQ_FEATURE	splice variant	0.913982 HSPB1, NFATC1	0.767939	1
Enrichment Score:				
Annotation Cluster 11	2.1054837611080806			
Category	Term	PValue	Genes	Fold Enrich Bonferroni
KEGG_PATHWAY	mmu05134:Legionellosis	0.00314	CXCL1, IL6, HSPA2, NFKBIA	13.16389 0.339742

	mmu04621:NOD-like receptor signaling pathway	0.00314 CXCL1, IL6, NFKBIA, TNFAIP3	13.16389	0.339742
KEGG_PATHWAY	mmu05164:Influenza A	0.011562 PRKCA, IL6, HSPA2, SOCS3, NFKBIA	5.484952	0.784574
KEGG_PATHWAY	mmu05162:Measles	0.033203 IL6, HSPA2, NFKBIA, TNFAIP3	5.517217	0.988405
Enrichment Score:				
Annotation Cluster 12	1.9600844654307423			
Category	Term	PValue	Genes	Fold Enrich Bonferroni
GO:0008285~negative regulation of cell proliferation				
GOTERM_BP_DIRECT	regulation of smooth muscle cell proliferation	0.001503 PRKCA, LIF, INHBA, IL6, FOSL1, FGF2	6.891006	0.67085
GO:0048661~positive regulation of ERK1 and ERK2 cascade				
GOTERM_BP_DIRECT	GO:0070374~positive regulation of ERK1 and ERK2 cascade	0.013524 PRKCA, IL6, FGF2	16.53841	0.999957
GOTERM_BP_DIRECT	cascade	0.064836 PRKCA, IL6, FGF2	7.037623	1
Enrichment Score:				
Annotation Cluster 13	1.7672878620366368			
Category	Term	PValue	Genes	Fold Enrich Bonferroni
mmu05161:Hepatitis B				
KEGG_PATHWAY	mmu05168:Herpes simplex infection	9.52E-05 PRKCA, FOS, IL6, MMP9, NFKBIA, CREB3L1, NFATC1	8.993819	0.012485
mmu04620:Toll-like receptor signaling pathway				
KEGG_PATHWAY	mmu05142:Chagas disease (American trypanosomiasis)	0.092929 FOS, IL6, SOCS3, NFKBIA	3.607411	0.999997
KEGG_PATHWAY		0.096586 FOS, IL6, NFKBIA	5.571843	0.999998
KEGG_PATHWAY		0.09983 FOS, IL6, NFKBIA	5.463651	0.999999
Enrichment Score:				
Annotation Cluster 14	1.6694518425935816			
Category	Term	PValue	Genes	Fold Enrich Bonferroni

KEGG_PATHWAY	mmu05161:Hepatitis B	9.52E-05	PRKCA, FOS, IL6, MMP9, NFKBIA, CREB3L1, NFATC1	8.993819	0.012485
KEGG_PATHWAY	mmu04380:Osteoclast differentiation	0.003978	FOS, SOCS3, NFKBIA, FOSL1, NFATC1	7.443864	0.409153
BIOCARTA	m_tcrPathway:T Cell Receptor Signaling Pathway	0.012678	PRKCA, FOS, NFKBIA, NFATC1	7.323864	0.690819
BIOCARTA	m_cdmacPathway:Cadmium induces DNA synthesis and proliferation in macrophages	0.013811	PRKCA, FOS, NFKBIA	15.10547	0.721814
KEGG_PATHWAY	mmu04921:Oxytocin signaling pathway	0.042485	PRKCA, FOS, CACNA2D1, NFATC1	5.002276	0.996755
KEGG_PATHWAY	mmu04662:B cell receptor signaling pathway	0.051019	FOS, NFKBIA, NFATC1	8.039373	0.999005
BIOCARTA	m_bcrPathway:BCR Signaling Pathway	0.057219	PRKCA, FOS, NFATC1	7.108456	0.995576
BIOCARTA	m_gpcrPathway:Signaling Pathway from G-Protein Families	0.057219	PRKCA, FOS, NFATC1	7.108456	0.995576
KEGG_PATHWAY	mmu04024:cAMP signaling pathway	0.081979	FOS, NFKBIA, CREB3L1, NFATC1	3.80884	0.999988
KEGG_PATHWAY	mmu04660:T cell receptor signaling pathway	0.096586	FOS, NFKBIA, NFATC1	5.571843	0.999998
BIOCARTA	m_keratinocytePathway:Keratinocyte Differentiation	0.116105	PRKCA, FOS, NFKBIA	4.738971	0.999988

Enrichment Score:

Annotation Cluster 15 1.4442820906558291

Category	Term	PValue	Genes	Fold Enrich	Bonferroni
KEGG_PATHWAY	mmu05161:Hepatitis B	9.52E-05	PRKCA, FOS, IL6, MMP9, NFKBIA, CREB3L1, NFATC1	8.993819	0.012485
KEGG_PATHWAY	mmu05031:Amphetamine addiction	0.047164	PRKCA, FOS, CREB3L1	8.399345	0.9983
KEGG_PATHWAY	mmu04725:Cholinergic synapse	0.116493	PRKCA, FOS, CREB3L1	4.980142	1

KEGG_PATHWAY	mmu04728:Dopaminergic synapse GO:0005783~endoplasmic reticulum	0.1535 PRKCA, FOS, CREB3L1 0.748245 PRKCA, FOS, CREB3L1	4.199672 1.114626	1 1
Enrichment Score:				
Annotation Cluster 16	1.335702433107032			
Category Term PValue Genes Fold Enrich Bonferroni				
UP_SEQ_FEATURE	metal ion-binding site:Calcium 3; via carbonyl oxygen	9.35E-04 PRKCA, MMP9, MMP3	64.32857	0.169803
UP_SEQ_FEATURE	metal ion-binding site:Calcium 3	0.002042 PRKCA, MMP9, MMP3	43.57742	0.334152
UP_SEQ_FEATURE	metal ion-binding site:Calcium 2; via carbonyl oxygen	0.0029 PRKCA, MMP9, MMP3	36.51081	0.438975
UP_SEQ_FEATURE	metal ion-binding site:Calcium 1	0.004644 PRKCA, MMP9, MMP3	28.74255	0.603965
UP_SEQ_FEATURE	metal ion-binding site:Calcium 2	0.006995 PRKCA, MMP9, MMP3	23.29138	0.752654
GOTERM_BP_DIRECT	GO:0006915~apoptotic process	0.036483 PRKCA, PIM1, NR4A1, TNFAIP3, DDIT4	3.868635	1
GOTERM_CC_DIRECT	GO:0043234~protein complex	0.127605 PRKCA, MMP9, NFKBIA, MMP3	3.130892	0.999986
UP_KEYWORDS	Calcium	0.178148 PRKCA, CACNA2D1, MMP9, MMP3	2.675554	1
GOTERM_MF_DIRECT	GO:0008270~zinc ion binding	0.231012 PRKCA, MMP9, NR4A1, TNFAIP3, MMP3 PRKCA, CACNA2D1, MMP9, PIM1, NR4A1, LIG4,	1.979126	1
UP_KEYWORDS	Metal-binding	0.391169 TNFAIP3, MMP3	1.303495	1
UP_KEYWORDS	Zinc	0.51349 PRKCA, MMP9, NR4A1, TNFAIP3, MMP3	1.317701	1
UP_KEYWORDS	Zinc-finger	0.773209 PRKCA, NR4A1, TNFAIP3 PRKCA, CACNA2D1, MMP9, NR4A1, LIG4, TNFAIP3,	1.060391	1
GOTERM_MF_DIRECT	GO:0046872~metal ion binding	0.808263 MMP3	0.887805	1

		Enrichment Score:			
Annotation Cluster 17	0.9401008508974922				
Category	Term	PValue	Genes	Fold Enrich	Bonferroni
GOTERM_BP_DIRECT	GO:0008283~cell proliferation	0.012607	PIM1, LIG4, HMGA2, DDIT4	8.018625	0.999915
UP_KEYWORDS	Cell cycle	0.302996	PIM1, LIG4, HMGA2	2.650978	1
GOTERM_BP_DIRECT	GO:0007049~cell cycle	0.395951	PIM1, LIG4, HMGA2	2.154842	1
		Enrichment Score:			
Annotation Cluster 18	0.534160811306344				
Category	Term	PValue	Genes	Fold Enrich	Bonferroni
			DUSP5, DUSP4, MMP9, DUSP10, PLA1A, TNFAIP3,		
UP_KEYWORDS	Hydrolase	0.066808	MMP3	2.352488	0.999007
GOTERM_MF_DIRECT	GO:0008270~zinc ion binding	0.231012	PRKCA, MMP9, NR4A1, TNFAIP3, MMP3	1.979126	1
UP_KEYWORDS	Protease	0.247788	MMP9, TNFAIP3, MMP3	3.061831	1
GOTERM_MF_DIRECT	GO:0016787~hydrolase activity	0.272762	DUSP4, MMP9, DUSP10, PLA1A, TNFAIP3, MMP3	1.66541	1
	GO:0008233~peptidase				
GOTERM_MF_DIRECT	activity	0.332293	MMP9, TNFAIP3, MMP3	2.473908	1
GOTERM_BP_DIRECT	GO:0006508~proteolysis	0.370491	MMP9, TNFAIP3, MMP3	2.273322	1
UP_KEYWORDS	Zinc	0.51349	PRKCA, MMP9, NR4A1, TNFAIP3, MMP3	1.317701	1
			PRKCA, CACNA2D1, MMP9, NR4A1, LIG4, TNFAIP3,		
GOTERM_MF_DIRECT	GO:0046872~metal ion binding	0.808263	MMP3	0.887805	1
		Enrichment Score:			
Annotation Cluster 19	0.3494247586609406				
Category	Term	PValue	Genes	Fold Enrich	Bonferroni
UP_KEYWORDS	Methylation	0.239062	DVL3, HSPA2, RIN1, HSPB1	2.304878	1
UP_KEYWORDS	Cytoskeleton	0.57048	HSPA2, RIN1, HSPB1	1.54661	1
GOTERM_CC_DIRECT	GO:0005856~cytoskeleton	0.656107	HSPA2, RIN1, HSPB1	1.324933	1
		Enrichment Score:			
Annotation Cluster 20	0.20317962059120928				
Category	Term	PValue	Genes	Fold Enrich	Bonferroni

UP_SEQ_FEATURE	binding site:ATP	0.361391 PRKCA, PIM1, LIG4	2.317153	1
UP_KEYWORDS	ATP-binding	0.43463 PRKCA, HSPA2, PIM1, LIG4	1.623392	1
UP_KEYWORDS	Nucleotide-binding	0.607193 PRKCA, HSPA2, PIM1, LIG4	1.261507	1
GOTERM_MF_DIRECT	GO:0005524~ATP binding	0.683485 PRKCA, HSPA2, PIM1, LIG4	1.129429	1
	GO:0000166~nucleotide			
GOTERM_MF_DIRECT	binding	0.945955 PRKCA, HSPA2, LIG4	0.659368	1
	GO:0005886~plasma			
GOTERM_CC_DIRECT	membrane	0.979263 PRKCA, PIM1, NFKBIA, RIN1, HSPB1, LIG4	0.605109	1
Enrichment Score:				
Annotation Cluster 21	7.100104747328335E-4			
Category	Term	PValue	Genes	Fold Enrich Bonferroni
	topological			
UP_SEQ_FEATURE	domain:Cytoplasmic	0.990623 CACNA2D1, CSF2RB, CREB3L1	0.469063	1
UP_SEQ_FEATURE	transmembrane region	0.999695 CACNA2D1, CSF2RB, CREB3L1	0.313288	1
UP_KEYWORDS	Membrane	0.999934 PRKCA, CACNA2D1, PIM1, RIN1, CSF2RB, CREB3L1	0.382244	1
UP_KEYWORDS	Transmembrane helix	0.999992 CACNA2D1, CSF2RB, CREB3L1	0.239192	1
UP_KEYWORDS	Transmembrane	0.999992 CACNA2D1, CSF2RB, CREB3L1	0.238607	1
	GO:0016021~integral			
GOTERM_CC_DIRECT	component of membrane	0.999999 CACNA2D1, CSF2RB, CREB3L1	0.214401	1

S. Table 2 Mouse GO IR compared to healthy fibroblasts_downregulated

Annotation Cluster 1	Enrichment Score:	PValue	Genes	Fold Enrich	Bonferroni
Category	Term				
UP_SEQ_FEATURE	propeptide:N-terminal propeptide	6.71E-08	COL3A1, COL1A2, COL1A1, COL11A1	414.069	1.11E-05
UP_SEQ_FEATURE	propeptide:C-terminal propeptide	1.17E-07	COL3A1, COL1A2, COL1A1, COL11A1	354.9163	1.93E-05
UP_SEQ_FEATURE	domain:Fibrillar collagen NC1	4.01E-07	COL3A1, COL1A2, COL1A1, COL11A1	248.4414	6.61E-05
INTERPRO	IPR000885:Fibrillar collagen, C-	4.11E-07	COL3A1, COL1A2, COL1A1, COL11A1	249.6242	4.65E-05
GOTERM_BP_DIRECT	GO:0030199~collagen fibril	4.22E-07	SFRP2, COL3A1, COL1A2, COL1A1, COL11A1	77.2735	2.61E-04
SMART	SM00038:COLFI	1.75E-06	COL3A1, COL1A2, COL1A1, COL11A1 VEGFB, BRAF, COL3A1, COL1A2, PDGFC, COL1A1,	151.6364	7.17E-05
KEGG_PATHWAY	mmu04510:Focal adhesion	3.94E-06	COL11A1, THBS4	11.00877	4.76E-04
KEGG_PATHWAY	mmu05146:Amoebiasis	3.81E-05	PLCB4, COL3A1, COL1A2, TLR4, COL1A1, COL11A1	14.60779	0.0046
	GO:0005201~extracellular matrix				
GOTERM_MF_DIRECT	structural constituent	4.22E-05	COL3A1, COL1A2, COL1A1, COL11A1	56.73496	0.005386
KEGG_PATHWAY	mmu04611:Platelet activation	6.55E-05	PLCB4, COL3A1, COL1A2, COL1A1, COL11A1, SYK	13.04665	0.007899
INTERPRO	IPR008160:Collagen triple helix	1.65E-04	COL3A1, COL1A2, COL1A1, COL11A1	36.12982	0.018441
UP_KEYWORDS	Collagen	1.73E-04	COL3A1, COL1A2, COL1A1, COL11A1	35.57647	0.018173
	GO:0048407~platelet-derived				
GOTERM_MF_DIRECT	growth factor binding	1.74E-04	COL3A1, COL1A2, COL1A1	145.3833	0.022063
UP_KEYWORDS	Hydroxylation	1.92E-04	COL3A1, COL1A2, COL1A1, COL11A1	34.36364	0.020119
KEGG_PATHWAY	mmu04512:ECM-receptor	1.97E-04	COL3A1, COL1A2, COL1A1, COL11A1, THBS4	16.18476	0.023603
UP_KEYWORDS	Extracellular matrix	2.18E-04	COL3A1, COL1A2, COL1A1, COL11A1, THBS4	16.08511	0.022813
GOTERM_CC_DIRECT	GO:0005581~collagen trimer	2.45E-04	COL3A1, COL1A2, COL1A1, COL11A1	31.58554	0.027529
UP_SEQ_FEATURE	region of interest:Triple-helical	3.51E-04	COL3A1, COL1A1, COL11A1	103.5172	0.056294
GOTERM_CC_DIRECT	GO:0031012~extracellular matrix	8.66E-04	COL3A1, COL1A2, COL1A1, COL11A1, THBS4 VEGFB, SFRP2, COL3A1, IL1RAP, COL1A2, PDGFC,	11.14626	0.094067
UP_KEYWORDS	Secreted	9.53E-04	COL1A1, COL11A1, THBS4	4.037982	0.096147
	GO:0005578~proteinaceous				
GOTERM_CC_DIRECT	extracellular matrix	0.001132	COL3A1, COL1A2, COL1A1, COL11A1, THBS4	10.37025	0.121142
	GO:0071230~cellular response to				
GOTERM_BP_DIRECT	amino acid stimulus	0.001346	COL3A1, COL1A2, PDGFC, COL1A1 VEGFB, SFRP2, COL3A1, IL1RAP, COL1A2, PDGFC,	17.59805	0.564964
GOTERM_CC_DIRECT	GO:0005576~extracellular region	0.003066	COL1A1, COL11A1, THBS4	3.36486	0.295372
KEGG_PATHWAY	mmu04974:Protein digestion and	0.003111	COL3A1, COL1A2, COL1A1, COL11A1	12.94781	0.314124

GOTERM_BP_DIRECT	GO:0001568~blood vessel	0.005763	COL3A1, COL1A2, COL1A1	25.46761	0.971896
GOTERM_BP_DIRECT	GO:0001501~skeletal system	0.012692	COL3A1, COL1A2, COL1A1	16.89907	0.999627
UP_KEYWORDS	Calcium	0.02019	COL3A1, COL1A2, COL1A1, COL11A1, THBS4 VEGFB, SFRP2, COL3A1, COL1A2, PDGFC, COL1A1,	4.570738	0.884911
GOTERM_CC_DIRECT	GO:0005615~extracellular space	0.02049	THBS4 ACVR2A, BRAF, COL3A1, COL1A2, COL1A1,	3.050399	0.905595
UP_KEYWORDS	Metal-binding	0.13205	COL11A1, RAD50, PHF6 ACVR2A, BRAF, COL3A1, COL1A2, COL1A1,	1.781443	1
GOTERM_MF_DIRECT	GO:0046872~metal ion binding	0.317813	COL11A1, RAD50, PHF6	1.386667	1
Enrichment Score:					
Annotation Cluster 2	3.242989941052708				
Category	Term	PValue	Genes	Fold Enrich	Bonferroni
GO:0050680~negative regulation					
GOTERM_BP_DIRECT	of epithelial cell proliferation	2.05E-04	SFRP2, RB1, GAS1, APC	33.48519	0.119192
GOTERM_BP_DIRECT	GO:0007050~cell cycle arrest	2.91E-04	RB1, GAS1, FOXO4, APC	29.76461	0.164579
GOTERM_BP_DIRECT	GO:0008285~negative regulation				
GOTERM_BP_DIRECT	of cell proliferation	0.003124	SFRP2, PBRM1, RB1, FOXO4, APC	7.84809	0.855414
Enrichment Score:					
Annotation Cluster 3	3.1525466853972954				
Category	Term	PValue	Genes	Fold Enrich	Bonferroni
KEGG_PATHWAY	mmu04151:PI3K-Akt signaling pathway	6.39E-09	VEGFB, KRAS, COL3A1, COL1A2, TLR4, PDGFC, GNG2, EFNA5, COL1A1, COL11A1, THBS4, SYK	9.738525	7.73E-07
KEGG_PATHWAY	mmu04510:Focal adhesion	3.94E-06	VEGFB, BRAF, COL3A1, COL1A2, PDGFC, COL1A1, COL11A1, THBS4	11.00877	4.76E-04
UP_KEYWORDS	Secreted	9.53E-04	VEGFB, SFRP2, COL3A1, IL1RAP, COL1A2, PDGFC, COL1A1, COL11A1, THBS4	4.037982	0.096147
UP_KEYWORDS	Disulfide bond	0.001049	VEGFB, ACVR2A, SFRP2, COL3A1, IL1RAP, COL1A2, TLR4, PDGFC, EFNA5, COL1A1, COL11A1, THBS4	2.903969	0.105281

UP_SEQ_FEATURE	signal peptide	COL3A1, TLR4, GAS1, VEGFB, ACVR2A, SFRP2, IL1RAP, COL1A2, EFNA5, PDGFC, COL1A1, COL3A1, EZH2, TLR4, GAS1, VEGFB, ACVR2A, IL1RAP, COL1A2, EFNA5, PDGFC, COL1A1,	0.00141	COL11A1, THBS4	2.584617	0.207704
UP_KEYWORDS	Glycoprotein	COL3A1, TLR4, GAS1, RB1, VEGFB, ACVR2A, SFRP2, IL1RAP, COL1A2, EFNA5, PDGFC, COL1A1, VEGFB, SFRP2, COL3A1, IL1RAP, COL1A2, PDGFC,	0.001548	COL11A1, THBS4	2.576147	0.1514
UP_KEYWORDS	Signal	VEGFB, SFRP2, COL3A1, COL1A2, PDGFC, COL1A1, VEGFB, ACVR2A, SFRP2, COL3A1, IL1RAP, TLR4, PDGFC, EFNA5, THBS4	0.002171	COL11A1, THBS4	2.329738	0.205752
GOTERM_CC_DIRECT	GO:0005576~extracellular region	VEGFB, SFRP2, COL3A1, COL1A2, PDGFC, COL1A1, VEGFB, ACVR2A, IL1RAP, COL1A2, TLR4, PDGFC, EFNA5, THBS4	0.003066	COL1A1, COL11A1, THBS4	3.36486	0.295372
GOTERM_CC_DIRECT	GO:0005615~extracellular space	ACVR2A, IL1RAP, COL1A2, TLR4, PDGFC, EFNA5, THBS4	0.02049	THBS4	3.050399	0.905595
UP_SEQ_FEATURE	disulfide bond	ACVR2A, IL1RAP, COL1A2, TLR4, PDGFC, EFNA5, THBS4	0.033027	PDGFC, EFNA5, THBS4	2.227064	0.996079
UP_SEQ_FEATURE	glycosylation site:N-linked (GlcNAc...)	VEGFB, SFRP2, COL3A1, COL1A2, PDGFC, COL1A1, VEGFB, ACVR2A, SFRP2, COL3A1, IL1RAP, TLR4, PDGFC, EFNA5, THBS4	0.084944	COL1A1, GAS1, COL11A1, THBS4	1.743204	1

Annotation Cluster 4	Enrichment Score: 2.5122576737300997					
	Category	Term	PValue	Genes	Fold Enrich	Bonferroni
	GOTERM_BP_DIRECT	GO:0051781~positive regulation of cell division	5.03E-05	VEGFB, PDGFC, THBS4, APC	53.5763	0.030608
	UP_KEYWORDS	Mitogen	9.68E-04	VEGFB, PDGFC, THBS4	63	0.097573
	UP_KEYWORDS	Growth factor	0.011442	VEGFB, PDGFC, THBS4 VEGFB, SFRP2, COL3A1, COL1A2, PDGFC, COL1A1,	17.85827	0.704729
	GOTERM_CC_DIRECT	GO:0005615~extracellular space	0.02049	THBS4	3.050399	0.905595
	GOTERM_MF_DIRECT	GO:0008083~growth factor	0.024053	VEGFB, PDGFC, THBS4	12.03172	0.955686

Annotation Cluster 5	Enrichment Score: 2.134573188535231					
	Category	Term	PValue	Genes	Fold Enrich	Bonferroni

KEGG_PATHWAY	mmu04014:Ras signaling pathway propeptide:Removed in mature	8.83E-04	VEGFB, KRAS, PDGFC, GNG2, EFNA5, FOXO4	7.463367	0.101389
UP_SEQ_FEATURE	form	0.005848	KRAS, GNG2, EFNA5, GAS1	10.43871	0.620052
UP_KEYWORDS	Lipoprotein	0.076431	KRAS, GNG2, EFNA5, GAS1	3.876923	0.999781
Enrichment Score:					
Annotation Cluster 6	2.1001117789576402				
Category	Term	PValue	Genes	Fold Enrich	Bonferroni
UP_KEYWORDS	Cell cycle	1.19E-04	RB1, GAS1, FOXO4, STAG2, RAD50, SMC3, HELLS	8.453674	0.012563
GOTERM_BP_DIRECT	GO:0007049~cell cycle GO:0000775~chromosome,	3.64E-04	RB1, GAS1, FOXO4, STAG2, RAD50, SMC3, HELLS	6.871553	0.201721
GOTERM_CC_DIRECT	centromeric region	0.001151	STAG2, SMC3, HELLS, PHF6	18.59291	0.123002
GOTERM_MF_DIRECT	GO:0003682~chromatin binding	0.00703	EZH2, PBRM1, STAG2, SMC3, HELLS	6.239628	0.59465
GOTERM_BP_DIRECT	GO:0051301~cell division	0.021545	RB1, STAG2, SMC3, HELLS	6.446346	0.999999
UP_KEYWORDS	Mitosis	0.043086	STAG2, SMC3, HELLS	8.756757	0.990613
KEGG_PATHWAY	mmu04110:Cell cycle GO:0007067~mitotic nuclear	0.065175	RB1, STAG2, SMC3	6.891577	0.999713
GOTERM_BP_DIRECT	division	0.072428	STAG2, SMC3, HELLS	6.527798	1
UP_KEYWORDS	Cell division	0.081513	STAG2, SMC3, HELLS	6.096774	0.999878
Enrichment Score:					
Annotation Cluster 7	2.062838212952619				
Category	Term	PValue	Genes	Fold Enrich	Bonferroni
GOTERM_CC_DIRECT	GO:0000775~chromosome, centromeric region	0.001151	STAG2, SMC3, HELLS, PHF6	18.59291	0.123002
GOTERM_CC_DIRECT	GO:0005694~chromosome	0.001548	EZH2, STAG2, RAD50, SMC3, PHF6	9.526163	0.16186
UP_KEYWORDS	Chromosome	0.001732	EZH2, STAG2, RAD50, SMC3, PHF6	9.242054	0.167868
GOTERM_MF_DIRECT	GO:0003682~chromatin binding	0.00703	EZH2, PBRM1, STAG2, SMC3, HELLS	6.239628	0.59465
UP_KEYWORDS	Centromere	0.012674	STAG2, SMC3, PHF6	16.92537	0.741272

GOTERM_CC_DIRECT	GO:0005634~nucleus	BRAF, EZH2, BNIP3, RB1, FOXO4, SMC3, RAD50, PLCB4, CASP12, PBRM1, PDGFC, STAG2, HELLS, SYK, APC, PHF6	0.0141	1.742216	0.801874
GOTERM_CC_DIRECT	GO:0005654~nucleoplasm	EZH2, PBRM1, BNIP3, FOXO4, STAG2, RAD50, SMC3, PHF6	0.019873	2.709664	0.898569
UP_KEYWORDS	Nucleus	BRAF, EZH2, PBRM1, PDGFC, RB1, FOXO4, STAG2, RAD50, SMC3, HELLS, PHF6	0.049023	1.834142	0.995147
UP_KEYWORDS	Phosphoprotein	BRAF, EZH2, BNIP3, RB1, FOXO4, SMC3, RAD50, CASP12, PBRM1, COL1A1, STAG2, HELLS, SYK, APC, PHF6	0.072031	1.488775	0.999638
Enrichment Score:					
Annotation Cluster 8	1.922270358680305				
Category	Term	PValue	Genes	Fold Enrich	Bonferroni
KEGG_PATHWAY	mmu04014:Ras signaling pathway	8.83E-04	VEGFB, KRAS, PDGFC, GNG2, EFNA5, FOXO4	7.463367	0.101389
UP_KEYWORDS	Developmental protein	0.007336	VEGFB, SFRP2, PDGFC, EFNA5, FOXO4, HELLS	4.647541	0.541815
GO:0007275~multicellular					
GOTERM_BP_DIRECT	organism development	0.022452	VEGFB, SFRP2, PDGFC, EFNA5, FOXO4, HELLS	3.51448	0.999999
GOTERM_BP_DIRECT	GO:0030154~cell differentiation	0.034637	VEGFB, BRAF, SFRP2, EFNA5, FOXO4	3.863675	1
UP_KEYWORDS	Differentiation	0.04857	VEGFB, SFRP2, EFNA5, FOXO4	4.681115	0.994895
Enrichment Score:					
Annotation Cluster 9	1.7765422608708206				
Category	Term	PValue	Genes	Fold Enrich	Bonferroni
KEGG_PATHWAY	mmu05218:Melanoma	0.001683	KRAS, BRAF, PDGFC, RB1	16.04799	0.18442
KEGG_PATHWAY	mmu05200:Pathways in cancer	0.001691	VEGFB, PLCB4, KRAS, BRAF, GNG2, RB1, APC	5.048008	0.185183
KEGG_PATHWAY	mmu05215:Prostate cancer	0.003111	KRAS, BRAF, PDGFC, RB1	12.94781	0.314124
KEGG_PATHWAY	mmu05219:Bladder cancer	0.008311	KRAS, BRAF, RB1	20.84282	0.635715
KEGG_PATHWAY	mmu05213:Endometrial cancer	0.013137	KRAS, BRAF, APC	16.43376	0.798128
mmu05223:Non-small cell lung					
KEGG_PATHWAY	cancer	0.015132	KRAS, BRAF, RB1	15.25992	0.841965
KEGG_PATHWAY	mmu05210:Colorectal cancer	0.019485	KRAS, BRAF, APC	13.35243	0.907534
KEGG_PATHWAY	mmu05214:Glioma	0.020062	KRAS, BRAF, RB1	13.14701	0.913893

KEGG_PATHWAY	mmu05212:Pancreatic cancer	0.020062 KRAS, BRAF, RB1	13.14701	0.913893
KEGG_PATHWAY	mmu05220:Chronic myeloid leukemia	0.0243 KRAS, BRAF, RB1	11.86883	0.949035
KEGG_PATHWAY	mmu04810:Regulation of actin cytoskeleton	0.034081 KRAS, BRAF, PDGFC, APC	5.349331	0.98494
UP_KEYWORDS	Methylation	0.122646 KRAS, BRAF, GNG2, RB1	3.15	0.999999
UP_KEYWORDS	Cytoplasm	0.689339 KRAS, BRAF, PDGFC, FOXO4, SYK, APC	1.029973	1

Enrichment Score:				
Annotation Cluster 10	1.7294883617400314	PValue	Genes	Fold Enrich Bonferroni
Category	Term			
KEGG_PATHWAY	mmu04015:Rap1 signaling pathway	6.49E-04	VEGFB, PLCB4, KRAS, BRAF, PDGFC, EFNA5	7.986501 0.075599
KEGG_PATHWAY	mmu05200:Pathways in cancer	0.001691	VEGFB, PLCB4, KRAS, BRAF, GNG2, RB1, APC	5.048008 0.185183
KEGG_PATHWAY	mmu04726:Serotonergic synapse	0.009628	PLCB4, KRAS, BRAF, GNG2	8.631874 0.689844
KEGG_PATHWAY	mmu04730:Long-term depression	0.017797	PLCB4, KRAS, BRAF	14.00911 0.886141
KEGG_PATHWAY	mmu04720:Long-term potentiation	0.020646	PLCB4, KRAS, BRAF	12.94781 0.919888
KEGG_PATHWAY	mmu04062:Chemokine signaling pathway	0.027542	PLCB4, KRAS, BRAF, GNG2	5.813303 0.965929
GOTERM_BP_DIRECT	GO:0007165~signal transduction	0.047147	KRAS, BRAF, IL1RAP, TLR4, GNG2, SMC3	2.881594 1
KEGG_PATHWAY	mmu04725:Cholinergic synapse	0.055309	PLCB4, KRAS, GNG2	7.562439 0.998977
UP_KEYWORDS	Methylation	0.122646	KRAS, BRAF, GNG2, RB1	3.15 0.999999
KEGG_PATHWAY	mmu05034:Alcoholism	0.148177	KRAS, BRAF, GNG2	4.230473 1

Enrichment Score:				
Annotation Cluster 11	1.4428988027085192	PValue	Genes	Fold Enrich Bonferroni
Category	Term			
GOTERM_BP_DIRECT	GO:0051276~chromosome organization	0.001685	RAD50, SMC3, APC	47.58421 0.647234
GOTERM_BP_DIRECT	GO:0006974~cellular response to DNA damage stimulus	0.145236	RAD50, SMC3, APC	4.305238 1

UP_KEYWORDS	Coiled coil	0.191751 PLCB4, PBRM1, STAG2, RAD50, SMC3, HELLS, APC	1.720416	1
Enrichment Score:				
Annotation Cluster 12	1.0575525256474256			
Category	Term	PValue	Genes	Fold Enrich Bonferroni
UP_KEYWORDS	Innate immunity	0.037822	IL1RAP, TLR4, SYK	9.410788 0.983209
UP_KEYWORDS	Immunity	0.092638	IL1RAP, TLR4, SYK	5.65586 0.999967
	GO:0002376~immune system			
GOTERM_BP_DIRECT	process	0.125109	IL1RAP, TLR4, SYK	4.721149 1
	GO:0045087~innate immune			
GOTERM_BP_DIRECT	response	0.134267	IL1RAP, TLR4, SYK	4.5205 1
Enrichment Score:				
Annotation Cluster 13	1.0419788437733186			
Category	Term	PValue	Genes	Fold Enrich Bonferroni
UP_KEYWORDS	Acetylation	0.00399	KRAS, BRAF, PBRM1, GNG2, RB1, FOXO4, STAG2,	2.66112 0.345467
UP_KEYWORDS	Chromatin regulator	0.046431	RAD50, SMC3, PHF6, APC	8.4 0.993524
UP_KEYWORDS	Nucleus	0.049023	EZH2, PBRM1, RB1, BRAF, EZH2, PBRM1, PDGFC, RB1, FOXO4, STAG2,	1.834142 0.995147
	GO:0016569~covalent chromatin		RAD50, SMC3, HELLS, PHF6	
GOTERM_BP_DIRECT	modification	0.067498	EZH2, PBRM1, RB1	6.797744 1
UP_KEYWORDS	Transcription regulation	0.075341	EZH2, PBRM1, RB1, FOXO4, HELLS, PHF6	2.521401 0.999752
UP_KEYWORDS	Transcription	0.08417	EZH2, PBRM1, RB1, FOXO4, HELLS, PHF6	2.440022 0.99991
GOTERM_MF_DIRECT	GO:0019899~enzyme binding	0.133177	RB1, FOXO4, PHF6	4.543229 1
	GO:0006351~transcription, DNA-			
GOTERM_BP_DIRECT	templated	0.178965	EZH2, PBRM1, RB1, FOXO4, HELLS, PHF6	1.918515 1
	GO:0006355~regulation of			
GOTERM_BP_DIRECT	transcription, DNA-templated	0.299104	EZH2, PBRM1, RB1, FOXO4, HELLS, PHF6	1.586836 1
UP_KEYWORDS	DNA-binding	0.33751	PBRM1, RB1, FOXO4, PHF6	1.885287 1
GOTERM_MF_DIRECT	GO:0003677~DNA binding	0.369165	PBRM1, RB1, FOXO4, RAD50, PHF6	1.574265 1

Enrichment Score:					
Annotation Cluster 14	0.9202226534608133	PValue	Genes	Fold Enrich	Bonferroni
Category	Term				
UP_KEYWORDS	Cell membrane	0.039925	GAS1, SYK, APC ACVR2A, KRAS, BRAF, IL1RAP, TLR4, PDGFC,	2.011173	0.986684
GOTERM_CC_DIRECT	GO:0005886~plasma membrane	0.081136	GNG2, EFNA5, GAS1, STAG2, SYK, APC BRAF, BNIP3, TLR4, GAS1, RAD50, VEGFB, ACVR2A, KRAS, IL1RAP, GNG2, EFNA5, PDGFC,	1.613623	0.999935
GOTERM_CC_DIRECT	GO:0016020~membrane	0.109983	STAG2, APC, SYK ACVR2A, KRAS, BRAF, IL1RAP, BNIP3, TLR4,	1.40483	0.999998
UP_KEYWORDS	Membrane	0.585233	PDGFC, GNG2, EFNA5, GAS1, SYK, APC	1.0448	1
Enrichment Score:					
Annotation Cluster 15	0.8645895536634423	PValue	Genes	Fold Enrich	Bonferroni
Category	Term				
UP_SEQ_FEATURE	nucleotide phosphate-binding region:ATP	0.015224	ACVR2A, BRAF, RAD50, SMC3, HELLS, SYK	3.869803	0.920439
UP_KEYWORDS	Nucleotide-binding	0.021528	ACVR2A, KRAS, BRAF, RAD50, SMC3, HELLS, SYK	3.017104	0.900426
UP_KEYWORDS	ATP-binding	0.027686	ACVR2A, BRAF, RAD50, SMC3, HELLS, SYK	3.327953	0.94901
KEGG_PATHWAY	mmu04650:Natural killer cell mediated cytotoxicity	0.048509	KRAS, BRAF, SYK	8.138624	0.997563
GOTERM_MF_DIRECT	GO:0000166~nucleotide binding	0.094989	ACVR2A, KRAS, BRAF, RAD50, SMC3, HELLS, SYK	2.102652	0.999997
GOTERM_MF_DIRECT	GO:0005524~ATP binding	0.100074	ACVR2A, BRAF, RAD50, SMC3, HELLS, SYK	2.315328	0.999999
INTERPRO	IPR027417:P-loop containing nucleoside triphosphate hydrolase	0.134397	KRAS, RAD50, SMC3, HELLS	3.020755	1
GOTERM_MF_DIRECT	GO:0004674~protein serine/threonine kinase activity	0.158419	ACVR2A, BRAF, SYK	4.076168	1
INTERPRO	IPR000719:Protein kinase, catalytic domain	0.163284	ACVR2A, BRAF, SYK	3.998835	1
UP_SEQ_FEATURE	domain:Protein kinase	0.182951	ACVR2A, BRAF, SYK	3.711774	1

		IPR011009:Protein kinase-like			
INTERPRO	domain	0.18393 ACVR2A, BRAF, SYK	3.703957	1	
	GO:0004672~protein kinase				
GOTERM_MF_DIRECT	activity	0.220436 ACVR2A, BRAF, SYK	3.285499	1	
UP_KEYWORDS	Kinase	0.228361 ACVR2A, BRAF, SYK	3.207921	1	
UP_SEQ_FEATURE	binding site:ATP	0.229176 ACVR2A, BRAF, SYK	3.196073	1	
	GO:0006468~protein				
GOTERM_BP_DIRECT	phosphorylation	0.235711 ACVR2A, BRAF, SYK	3.139236	1	
GOTERM_BP_DIRECT	GO:0016310~phosphorylation	0.257275 ACVR2A, BRAF, SYK	2.954575	1	
UP_SEQ_FEATURE	active site:Proton acceptor	0.303073 ACVR2A, BRAF, SYK	2.624381	1	
GOTERM_MF_DIRECT	GO:0016301~kinase activity	0.30928 ACVR2A, BRAF, SYK	2.588427	1	
UP_KEYWORDS	Transferase	0.355473 ACVR2A, BRAF, EZH2, SYK	1.828295	1	
GOTERM_MF_DIRECT	GO:0016740~transferase activity	0.447634 ACVR2A, BRAF, EZH2, SYK	1.580254	1	

Enrichment Score:

Annotation Cluster 16 0.021827864066571737

Category	Term	PValue	Genes	Fold Enrich	Bonferroni
UP_KEYWORDS	Receptor	0.862946	ACVR2A, IL1RAP, TLR4	0.867968	1
UP_SEQ_FEATURE	topological domain:Extracellular	0.882021	ACVR2A, IL1RAP, TLR4	0.825935	1
UP_SEQ_FEATURE	topological domain:Cytoplasmic	0.951958	ACVR2A, IL1RAP, TLR4	0.646983	1
UP_SEQ_FEATURE	transmembrane region	0.977845	ACVR2A, IL1RAP, BNIP3, TLR4	0.576163	1
	GO:0016021~integral component				
GOTERM_CC_DIRECT	of membrane	0.997353	ACVR2A, SFRP2, IL1RAP, BNIP3, TLR4	0.476447	1
UP_KEYWORDS	Transmembrane helix	0.997678	ACVR2A, IL1RAP, BNIP3, TLR4	0.43586	1
UP_KEYWORDS	Transmembrane	0.997735	ACVR2A, IL1RAP, BNIP3, TLR4	0.434795	1

S. Table 2 Mouse GO IR compared to healthy fibroblasts_upregulated

Enrichment Score:					
Annotation Cluster 1	Term	PValue	Genes	Fold Enrich	Bonferroni
Category	Growth factor	8.99E-07	CXCL1, INHBA, IL6, NGF, IL11 CXCL1, INHBA, IL6, CLCF1, FST, PLA1A, WNT11,	59.52756	4.94E-05
UP_KEYWORDS	Secreted	1.83E-06	NGF, IL11	8.075964	1.00E-04
GOTERM_MF_DIRECT	GO:0008083~growth factor activity	4.30E-06	CXCL1, INHBA, IL6, NGF, IL11 CXCL1, INHBA, IL6, CLCF1, FST, PLA1A, WNT11,	40.10575	2.62E-04
GOTERM_CC_DIRECT	GO:0005576~extracellular region	7.22E-06	NGF, IL11	6.72972	1.88E-04
GOTERM_MF_DIRECT	GO:0005125~cytokine activity	2.00E-05	CXCL1, INHBA, IL6, CLCF1, IL11	27.17445	0.00122
UP_KEYWORDS	Cytokine	2.29E-04	CXCL1, IL6, CLCF1, IL11	30.24	0.012511
GOTERM_MF_DIRECT	GO:0005102~receptor binding	2.54E-04	INHBA, IL6, CLCF1, WNT11, NGF	14.11489	0.015392
GOTERM_CC_DIRECT	GO:0005615~extracellular space mmu04060:Cytokine-cytokine receptor interaction	3.48E-04	CXCL1, INHBA, IL6, CLCF1, WNT11, NGF, IL11	6.100798	0.009013
KEGG_PATHWAY		7.46E-04	CXCL1, INHBA, IL6, CLCF1, IL11 CXCL1, INHBA, IL6, CLCF1, FST, PLA1A, WNT11,	10.59366	0.038789
UP_SEQ_FEATURE	signal peptide	9.02E-04	NGF, IL11	3.459411	0.049253
INTERPRO	IPR009079:Four-helical cytokine-like, core	0.001698	IL6, CLCF1, IL11 CXCL1, INHBA, IL6, CLCF1, FST, PLA1A, WNT11,	45.26154	0.081447
UP_KEYWORDS	Signal	0.002414	NGF, IL11	2.995378	0.12449
UP_KEYWORDS	Disulfide bond	0.00757	CXCL1, INHBA, IL6, FST, PLA1A, WNT11, NGF	3.387964	0.341603
GOTERM_BP_DIRECT	GO:0008284~positive regulation of cell proliferation	0.007618	IL6, CLCF1, NGF, IL11	8.896433	0.917953
UP_SEQ_FEATURE	disulfide bond	0.035235	CXCL1, INHBA, IL6, FST, PLA1A, NGF	2.870438	0.865842
UP_KEYWORDS	Glycoprotein	0.071248	INHBA, CLCF1, FST, PLA1A, WNT11, NGF	2.377982	0.982841
UP_SEQ_FEATURE	glycosylation site:N-linked (GlcNAc...)	0.125099	INHBA, CLCF1, FST, PLA1A, WNT11, NGF	2.022116	0.999438
GOTERM_BP_DIRECT	GO:0045944~positive regulation of transcription from RNA polymerase II promoter	0.178087	INHBA, IL6, IL11	3.634573	1

	Enrichment Score:			
Annotation Cluster 2	2.858676753889115			
Category	Term	PValue	Genes	Fold Enrich Bonferroni
GOTERM_MF_DIRECT	GO:0005102~receptor binding	2.54E-04	INHBA, IL6, CLCF1, WNT11, NGF	14.11489 0.015392
GOTERM_CC_DIRECT	GO:0005615~extracellular space	3.48E-04	CXCL1, INHBA, IL6, CLCF1, WNT11, NGF, IL11	6.100798 0.009013
	GO:0043065~positive regulation of			
GOTERM_BP_DIRECT	apoptotic process	0.001971	IL6, GADD45G, WNT11, NGF	14.39363 0.47544
	GO:0010628~positive regulation of			
GOTERM_BP_DIRECT	gene expression	0.003239	INHBA, IL6, WNT11, NGF	12.08488 0.65382
	GO:0045893~positive regulation of			
GOTERM_BP_DIRECT	transcription, DNA-templated	0.009004	INHBA, IL6, PIM1, WNT11	8.371296 0.948061
	Enrichment Score:			
Annotation Cluster 3	1.891029378739467			
Category	Term	PValue	Genes	Fold Enrich Bonferroni
KEGG_PATHWAY	mmu04621:NOD-like receptor signaling pathway	0.004638	CXCL1, IL6, TNFAIP3	26.98596 0.218389
KEGG_PATHWAY	mmu04668:TNF signaling pathway	0.016207	CXCL1, IL6, TNFAIP3	14.11193 0.579369
	GO:0006954~inflammatory			
GOTERM_BP_DIRECT	response	0.02824	CXCL1, IL6, TNFAIP3	10.51279 0.999915
	Enrichment Score:			
Annotation Cluster 4	0.9077843484664367			
Category	Term	PValue	Genes	Fold Enrich Bonferroni
UP_KEYWORDS	Apoptosis	0.035572	GADD45G, PIM1, TNFAIP3	9.276074 0.863589
GOTERM_BP_DIRECT	GO:0006915~apoptotic process	0.070289	GADD45G, PIM1, TNFAIP3	6.344561 1
GOTERM_MF_DIRECT	GO:0005515~protein binding	0.214551	IL6, CLCF1, GADD45G, PIM1, WNT11, TNFAIP3	1.705376 1
GOTERM_CC_DIRECT	GO:0005634~nucleus	0.435854	DUSP4, FST, GADD45G, PIM1, DUSP10, TNFAIP3	1.306662 1
	Enrichment Score:			
Annotation Cluster 5	0.7673855924904919			

Category	Term	PValue	Genes	Fold Enrich	Bonferroni
UP_KEYWORDS	Hydrolase	0.076071	DUSP4, DUSP10, PLA1A, TNFAIP3	3.674362	0.987114
GOTERM_MF_DIRECT	GO:0016787~hydrolase activity	0.118836	DUSP4, DUSP10, PLA1A, TNFAIP3	3.034747	0.999555
UP_KEYWORDS	Nucleus	0.551668	DUSP4, PIM1, DUSP10, TNFAIP3	1.333921	1
Enrichment Score:					
Annotation Cluster 6	0.19357586408446692				
Category	Term	PValue	Genes	Fold Enrich	Bonferroni
UP_KEYWORDS	Cytoplasm	0.529924	PIM1, DUSP10, RIN1, TNFAIP3	1.373297	1
UP_KEYWORDS	Nucleus	0.551668	DUSP4, PIM1, DUSP10, TNFAIP3	1.333921	1
UP_KEYWORDS	Phosphoprotein	0.898217	DUSP4, PIM1, RIN1, TNFAIP3	0.794013	1

S. Table 3 Mouse GO TH17 compared to IR_upregulated

Annotation Cluster 1	Enrichment Score: 5.38189138476703	PValue	Genes	Fold Enrich	Bonferroni
Category	Term GO:0051781~positive regulation of cell division				
GOTERM_BP_DIRECT	cell division	5.32E-08	PGF, IL1B, PDGFC, IL1A, THBS4, APC	57.40317	4.31E-05
UP_KEYWORDS	Mitogen GO:0045766~positive regulation of angiogenesis	5.19E-07	PGF, IL1B, PDGFC, IL1A, THBS4	75	6.38E-05
GOTERM_BP_DIRECT	angiogenesis	0.002589	PGF, MMP9, IL1B, IL1A	14.23219	0.877473
Enrichment Score:					
Annotation Cluster 2	5.282045892666902				
Category	Term	PValue	Genes	Fold Enrich	Bonferroni
UP_KEYWORDS	Growth factor	2.80E-09	CXCL1, LIF, CSF2, IL6, PGF, PDGFC, BMP5, THBS4	34.01575	3.45E-07
			CXCL1, CSF2, IL6, TNF, PGF, MMP9, MMP3, LIF, IL1RAP, IL1B, PDGFC, COL1A1, IL1A, BMP5, CD14,		
UP_KEYWORDS	Secreted	1.39E-08	THBS4, SPP1	5.448071	1.71E-06
			CXCL1, CSF2, IL6, TNF, PGF, MMP9, MMP3, LIF, HSP90B1, IL1RAP, IL1B, PDGFC, COL1A1, IL1A,		
GOTERM_CC_DIRECT	GO:0005576~extracellular region	2.49E-08	BMP5, CD14, THBS4, SPP1	4.806943	2.82E-06
GOTERM_MF_DIRECT	GO:0008083~growth factor activity	4.20E-08	CXCL1, LIF, CSF2, IL6, PGF, PDGFC, BMP5, THBS4	22.91757	5.47E-06
			CXCL1, CSF2, IL6, TNF, PGF, MMP9, MMP3, LIF,		
GOTERM_CC_DIRECT	GO:0005615~extracellular space	1.58E-07	IL1B, PDGFC, COL1A1, IL1A, BMP5, CD14, THBS4,	4.980243	1.79E-05
			CXCL1, CSF2, IL6, TNF, PGF, MMP9, TLR4, MMP3,		
UP_KEYWORDS	Disulfide bond	5.42E-05	LIF, HSP90B1, IL1RAP, PDGFC, COL1A1, BMP5,		
			CD14, THBS4, CSF1R	2.93854	0.006642
UP_SEQ_FEATURE	disulfide bond	1.78E-04	CXCL1, CSF2, IL6, TNF, PGF, MMP9, TLR4, MMP3,		
			LIF, HSP90B1, IL1RAP, PDGFC, BMP5, CD14, THBS4,		
UP_KEYWORDS	Glycoprotein	5.87E-04	CSF1R, SPP1	2.406291	0.069685
			CXCL1, CSF2, IL6, PGF, MMP9, TLR4, MMP3, LIF,		
UP_SEQ_FEATURE	signal peptide	6.01E-04	HSP90B1, IL1RAP, PDGFC, COL1A1, BMP5, CD14,		
			THBS4, CSF1R, SPP1	2.39065	0.123903

UP_KEYWORDS	Signal	0.001391	CXCL1, CSF2, IL6, PGF, MMP9, TLR4, MMP3, LIF, HSP90B1, ENDOG, IL1RAP, PDGFC, COL1A1, BMP5, CD14, THBS4, CSF1R, SPP1	2.139555	0.15732
UP_SEQ_FEATURE	glycosylation site:N-linked (GlcNAc...)	0.002584	CSF1R, SPP1	2.096096	0.433995
Annotation Cluster 3	Enrichment Score: 3.6147870774603135				
Category	Term	PValue	Genes	Fold Enrich	Bonferroni
KEGG_PATHWAY	mmu04668:TNF signaling pathway	1.28E-10	CXCL1, LIF, FOS, CSF2, IL6, CASP3, TNF, SOCS3, MMP9, IL1B, MMP3	18.93063	1.73E-08
UP_KEYWORDS	Cytokine	2.37E-09	CXCL1, LIF, CSF2, IL6, TNF, IL1B, BMP5, IL1A, SPP1	24.3	2.91E-07
GOTERM_BP_DIRECT	GO:0006954~inflammatory response	3.62E-09	CXCL1, LIF, FOS, CSF2, IL6, IL1A, SPP1	13.76675	2.93E-06
KEGG_PATHWAY	mmu05134:Legionellosis	1.30E-08	CXCL1, IL6, CASP3, TNF, IL1B, BNIP3, TLR4, CD14	26.32777	1.75E-06
GOTERM_MF_DIRECT	GO:0005125~cytokine activity	3.03E-08	CXCL1, LIF, CSF2, IL6, TNF, IL1B, BMP5, IL1A, SPP1	17.46929	3.94E-06
KEGG_PATHWAY	mmu05133:Pertussis	8.27E-08	FOS, IL6, CASP3, TNF, IL1B, TLR4, CD14, IL1A	20.2795	1.12E-05
KEGG_PATHWAY	mmu05132:Salmonella infection	1.20E-07	CXCL1, FOS, CSF2, IL6, IL1B, TLR4, CD14, IL1A	19.23952	1.62E-05
	GO:0008285~negative regulation of		LIF, IL6, TNF, CDKN2C, IL1B, PBRM1, BMP5, IL1A,		
GOTERM_BP_DIRECT	cell proliferation	1.54E-07	CSF1R, APC	11.21156	1.25E-04
KEGG_PATHWAY	mmu05323:Rheumatoid arthritis	1.70E-07	FOS, CSF2, IL6, TNF, IL1B, TLR4, MMP3, IL1A	18.30101	2.29E-05
GOTERM_BP_DIRECT	GO:0019221~cytokine-mediated	8.90E-07	IL6, KRAS, SOCS3, IL1RAP, IL1B, IL1A, CSF1R	20.64155	7.21E-04
KEGG_PATHWAY	mmu05146:Amoebiasis	1.94E-06	CSF2, IL6, CASP3, TNF, IL1B, TLR4, COL1A1, CD14	12.82635	2.62E-04
KEGG_PATHWAY	mmu04640:Hematopoietic cell lineage	4.33E-06	CSF2, IL6, TNF, IL1B, CD14, IL1A, CSF1R	15.44821	5.85E-04
GOTERM_BP_DIRECT	GO:0032755~positive regulation of	5.69E-06	IL6, TNF, IL1B, TLR4, IL1A	41.39652	0.0046
	interleukin-6 production				
	mmu04620:Toll-like receptor signaling				
KEGG_PATHWAY	pathway	1.18E-05	FOS, IL6, TNF, IL1B, TLR4, CD14, SPP1	13.00097	0.001593
GOTERM_BP_DIRECT	GO:0046330~positive regulation of	1.87E-05	TNF, IL1B, TLR4, GADD45A, IL1A	30.7517	0.015047
	JNK cascade				

KEGG_PATHWAY	mmu04060:Cytokine-cytokine receptor interaction GO:0010628~positive regulation of gene expression	2.75E-05 CXCL1, LIF, CSF2, IL6, TNF, IL1RAP, IL1B, IL1A, CSF1R	6.976315	0.003707
GOTERM_BP_DIRECT	GO:0006955~immune response	2.84E-05 LIF, CSF2, IL6, KRAS, TNF, IL1B, TLR4, IL1A	8.632056	0.022767
GOTERM_BP_DIRECT	GO:0031663~lipopolysaccharide-mediated signaling pathway	3.17E-05 CXCL1, LIF, CSF2, IL6, TNF, IL1B, IL1A	11.07966	0.025338
GOTERM_BP_DIRECT	GO:0070374~positive regulation of ERK1 and ERK2 cascade	4.65E-05 TNF, IL1B, TLR4, CD14	55.55146	0.036995
GOTERM_BP_DIRECT	mmu05144:Malaria	6.37E-05 IL6, IL1B, TLR4, PDGFC, IL1A, CSF1R	13.74012	0.050301
KEGG_PATHWAY	GO:0051092~positive regulation of NF-kappaB transcription factor activity	1.04E-04 IL6, TNF, IL1B, TLR4, THBS4	19.54014	0.013877
GOTERM_BP_DIRECT	GO:0045429~positive regulation of nitric oxide biosynthetic process	1.07E-04 KRAS, TNF, IL1RAP, IL1B, TLR4	19.7488	0.082689
GOTERM_BP_DIRECT	GO:0032729~positive regulation of interferon-gamma production	1.44E-04 IL6, TNF, IL1B, TLR4	38.26878	0.109892
GOTERM_BP_DIRECT	mmu05321:Inflammatory bowel disease (IBD)	2.21E-04 TNF, IL1B, TLR4, CD14	33.11722	0.164167
KEGG_PATHWAY	mmu05152:Tuberculosis	2.32E-04 IL6, TNF, IL1B, TLR4, IL1A	15.89706	0.030882
KEGG_PATHWAY	mmu05140:Leishmaniasis	2.65E-04 IL6, CASP3, TNF, IL1B, TLR4, CD14, IL1A	7.460782	0.035172
KEGG_PATHWAY	mmu04380:Osteoclast differentiation	3.18E-04 FOS, TNF, IL1B, TLR4, IL1A	14.65511	0.042072
KEGG_PATHWAY	mmu05020:Prion diseases	4.53E-04 FOS, TNF, SOCS3, IL1B, IL1A, CSF1R	8.932636	0.059334
KEGG_PATHWAY	GO:0005149~interleukin-1 receptor binding	6.38E-04 IL6, CASP12, IL1B, IL1A	22.73762	0.08258
GOTERM_MF_DIRECT	GO:0042346~positive regulation of NF-kappaB import into nucleus	7.17E-04 IL1RAP, IL1B, IL1A	73.30252	0.088987
GOTERM_BP_DIRECT	GO:0032722~positive regulation of chemokine production	8.37E-04 TNF, IL1B, TLR4	67.97744	0.492523
GOTERM_BP_DIRECT	GO:0032874~positive regulation of stress-activated MAPK cascade	9.29E-04 IL6, TNF, TLR4	64.57857	0.528872
GOTERM_BP_DIRECT	GO:0032496~response to lipopolysaccharide	9.29E-04 IL1B, TLR4, IL1A	64.57857	0.528872
GOTERM_BP_DIRECT	mmu04932:Non-alcoholic fatty liver disease (NAFLD)	0.00101 CXCL1, FOS, TNF, IL1B, TLR4	10.927	0.558786
KEGG_PATHWAY		0.001229 IL6, CASP3, TNF, SOCS3, IL1B, IL1A	7.168867	0.152965

GOTERM_BP_DIRECT	GO:0071222~cellular response to lipopolysaccharide	0.001256 CSF2, IL6, TNF, TLR4, CD14	10.29961	0.638829
GOTERM_BP_DIRECT	GO:0032757~positive regulation of interleukin-8 production	0.001456 TNF, IL1B, TLR4	51.66286	0.692774
KEGG_PATHWAY	mmu04064:NF-kappa B signaling pathway	0.001535 TNF, BCL2A1A, IL1B, TLR4, CD14	9.669349	0.18724
GOTERM_BP_DIRECT	GO:0045944~positive regulation of transcription from RNA polymerase II promoter	0.001541 LIF, FOS, IL6, TNF, ETS2, IL1B, TLR4, BMP5, IL1A	3.894185	0.713167
KEGG_PATHWAY	mmu05164:Influenza A	0.001796 IL6, TNF, SOCS3, IL1B, TLR4, IL1A	6.581943	0.215448
GOTERM_BP_DIRECT	GO:0051091~positive regulation of sequence-specific DNA binding	0.001824 IL6, TNF, IL1B, TLR4	16.09435	0.772023
BIOCARTA	M_il1rPathway:Signal transduction through IL1R	0.001884 IL6, TNF, IL1RAP, IL1B, IL1A	8.491436	0.131899
KEGG_PATHWAY	mmu05142:Chagas disease (American trypanosomiasis)	0.001915 FOS, IL6, TNF, IL1B, TLR4	9.106086	0.227975
GOTERM_BP_DIRECT	GO:0045840~positive regulation of mitotic nuclear division	0.001959 TNF, IL1B, IL1A	44.53695	0.795754
GOTERM_CC_DIRECT	GO:0045121~membrane raft	0.002118 CASP3, KRAS, TNF, TLR4, CD14	8.934024	0.21307
KEGG_PATHWAY	mmu05332:Graft-versus-host disease	0.002415 IL6, TNF, IL1B, IL1A	14.42964	0.278452
GOTERM_BP_DIRECT	GO:2001240~negative regulation of extrinsic apoptotic signaling pathway	0.002533 CSF2, TNF, IL1B	39.13853	0.871846
GOTERM_BP_DIRECT	in absence of ligand	0.002688 TNF, CD14, IL1A	37.98739	0.886955
KEGG_PATHWAY	mmu04621:NOD-like receptor	0.00314 CXCL1, IL6, TNF, IL1B	13.16389	0.345942
KEGG_PATHWAY	signaling pathway	0.003785 CASP3, KRAS, TNF, MMP9, TLR4, COL1A1	5.544395	0.400669
KEGG_PATHWAY	mmu05168:Herpes simplex infection	0.004199 FOS, IL6, CASP3, TNF, SOCS3, IL1B	5.411116	0.433403
GOTERM_BP_DIRECT	GO:0043123~positive regulation of I-	0.004648 TNF, IL1B, TLR4, IL1A	11.55769	0.977036
KEGG_PATHWAY	kappaB kinase/NF-kappaB signaling	0.005221 CCNE2, IL6, IL1B, TLR4, IL1A	6.896521	0.506723
	mmu05162:Measles			

	m_inflamPathway:Cytokines and Inflammatory Response	0.008705 CSF2, IL6, TNF, IL1A	8.622074	0.480949
GOTERM_CC_DIRECT	GO:0009986~cell surface	0.009545 TNF, TLR4, PDGFC, CD14, IL1A, CSF1R	4.465592	0.661697
GOTERM_BP_DIRECT	GO:0043525~positive regulation of neuron apoptotic process	0.010691 CASP3, TNF, IL1B	18.71843	0.999834
GOTERM_BP_DIRECT	GO:0000187~activation of MAPK activity	0.012859 TNF, IL1B, TLR4	16.99436	0.999972
KEGG_PATHWAY	mmu05143:African trypanosomiasis	0.014082 IL6, TNF, IL1B	16.07875	0.852606
BIOCARTA	m_il10Pathway:IL-10 Anti-inflammatory Signaling Pathway	0.016555 IL6, TNF, IL1A	14.01087	0.714083
BIOCARTA	m_hsp27Pathway:Stress Induction of HSP Regulation	0.02236 CASP3, TNF, IL1A	12.00932	0.816586
BIOCARTA	m_erythPathway:Erythrocyte Differentiation Pathway	0.025535 CSF2, IL6, IL1A	11.2087	0.856294
GOTERM_CC_DIRECT	GO:0009897~external side of plasma membrane	0.028187 IL6, TNF, TLR4, CD14	5.907165	0.960479
KEGG_PATHWAY	mmu04940:Type I diabetes mellitus	0.040995 TNF, IL1B, IL1A	9.076711	0.996486
BIOCARTA	m_cytokinePathway:Cytokine Network	0.048012 IL6, TNF, IL1A	8.006211	0.975034
BIOCARTA	m_nfkbpathway:NF-κB Signaling Pathway	0.061191 TNF, TLR4, IL1A	7.005435	0.991224
KEGG_PATHWAY	mmu05010:Alzheimer's disease	0.063668 CASP3, TNF, CASP12, IL1B	4.239217	0.999861
KEGG_PATHWAY	mmu05145:Toxoplasmosis	0.103104 CASP3, TNF, TLR4	5.359582	1
KEGG_PATHWAY	mmu04931:Insulin resistance	0.11142 IL6, TNF, SOCS3	5.115965	1

Enrichment Score: Annotation Cluster 4				
Category	Term	PValue	Genes	Fold Enrich Bonferroni
GOTERM_BP_DIRECT	GO:0008284~positive regulation of cell proliferation	2.47E-05	CSF1R LIF, CSF2, PLA2G4A, IL6, KRAS, TNF, PGF, PDGFC,	7.148919 0.019815
KEGG_PATHWAY	mmu04014:Ras signaling pathway	0.006301	PLA2G4A, KRAS, PGF, ETS2, PDGFC, CSF1R	4.9149 0.573993
KEGG_PATHWAY	mmu04015:Rap1 signaling pathway	0.099149	KRAS, PGF, PDGFC, CSF1R	3.506269 0.999999

	Enrichment Score:			
Annotation Cluster 5	2.4083097272040006			
Category	Term	PValue	Genes	Fold Enrich Bonferroni
GOTERM_BP_DIRECT	GO:0006955~immune response	3.17E-05	CXCL1, LIF, CSF2, IL6, TNF, IL1B, IL1A	11.07966 0.025338
INTERPRO	IPR012351:Four-helical cytokine, core	0.013287	LIF, CSF2, IL6	16.71591 0.894296
	IPR009079:Four-helical cytokine-like,			
INTERPRO	core	0.01416	LIF, CSF2, IL6	16.16484 0.908911
	mmu04630:Jak-STAT signaling pathway			
KEGG_PATHWAY	pathway	0.039033	LIF, CSF2, IL6, SOCS3	5.174769 0.995369
	Enrichment Score:			
Annotation Cluster 6	1.9743105321604364			
Category	Term	PValue	Genes	Fold Enrich Bonferroni
GOTERM_CC_DIRECT	GO:0031012~extracellular matrix	0.00321	HSP90B1, MMP9, COL1A1, MMP3, THBS4	7.961613 0.304601
UP_KEYWORDS	Extracellular matrix	0.008761	MMP9, COL1A1, MMP3, THBS4	9.191489 0.661216
UP_KEYWORDS	Calcium	0.016114	PLA2G4A, HSP90B1, MMP9, COL1A1, MMP3, THBS4	3.917775 0.864411
	GO:0005578~proteinaceous extracellular matrix			
GOTERM_CC_DIRECT	extracellular matrix	0.02796	MMP9, COL1A1, MMP3, THBS4	5.925859 0.959422
	Enrichment Score:			
Annotation Cluster 7	1.9577726206341322			
Category	Term	PValue	Genes	Fold Enrich Bonferroni
	mmu04664:Fc epsilon RI signaling pathway	0.00517	CSF2, PLA2G4A, KRAS, TNF	11.03443 0.503302
KEGG_PATHWAY	mmu04660:T cell receptor signaling pathway	0.01528	FOS, CSF2, KRAS, TNF	7.429123 0.874916
KEGG_PATHWAY	mmu04650:Natural killer cell mediated cytotoxicity	0.016945	CSF2, CASP3, KRAS, TNF	7.146109 0.900463
	Enrichment Score:			
Annotation Cluster 8	1.8334503554017587			
Category	Term	PValue	Genes	Fold Enrich Bonferroni
GOTERM_MF_DIRECT	GO:0002020~protease binding	0.002603	CASP3, TNF, CASP12, APC	14.20106 0.287371
KEGG_PATHWAY	mmu04210:Apoptosis	0.003633	CASP3, TNF, ENDOG, CASP12	12.50569 0.388191

KEGG_PATHWAY	mmu05014:Amyotrophic lateral sclerosis (ALS)	0.028644 CASP3, TNF, CASP12	11.03443	0.980228
GOTERM_BP_DIRECT	GO:0006915~apoptotic process	0.039455 CASP3, TNF, BCL2A1A, CASP12, BNIP3	3.776525	1
KEGG_PATHWAY	mmu05010:Alzheimer's disease	0.063668 CASP3, TNF, CASP12, IL1B	4.239217	0.999861
Enrichment Score:				
Annotation Cluster 9	1.7241640153216549	PValue Genes	Fold Enrich	Bonferroni
Category	Term			
KEGG_PATHWAY	mmu04110:Cell cycle	0.003757 CCNE2, CDKN2C, GADD45A, STAG2, SMC3	7.563926	0.398371
UP_KEYWORDS	Cell cycle	0.005199 CCNE2, CDKN2C, LIG4, GADD45A, STAG2, SMC3	5.175719	0.473307
GOTERM_BP_DIRECT	GO:0007049~cell cycle	0.012113 CCNE2, CDKN2C, LIG4, GADD45A, STAG2, SMC3	4.207073	0.999949
UP_KEYWORDS	Cell division	0.029474 CCNE2, LIG4, STAG2, SMC3	5.806452	0.974773
GOTERM_BP_DIRECT	GO:0051301~cell division	0.052508 CCNE2, LIG4, STAG2, SMC3	4.604533	1
UP_KEYWORDS	DNA damage	0.123241 LIG4, GADD45A, SMC3	4.821429	1
Enrichment Score:				
Annotation Cluster 10	1.613965559981692	PValue Genes	Fold Enrich	Bonferroni
Category	Term			
KEGG_PATHWAY	mmu05205:Proteoglycans in cancer	0.003785 CASP3, KRAS, TNF, MMP9, TLR4, COL1A1	5.544395	0.400669
UP_KEYWORDS	Polymorphism	0.054808 TNF, MMP9, TLR4	7.751196	0.999025
UP_SEQ_FEATURE	sequence variant	0.069374 TNF, MMP9, TLR4, APC	4.096196	1
Enrichment Score:				
Annotation Cluster 11	1.5512630368546512	PValue Genes	Fold Enrich	Bonferroni
Category	Term			
	short sequence motif:Cell attachment			
UP_SEQ_FEATURE	site	0.013301 COL1A1, THBS4, SPP1	16.68293	0.947445
KEGG_PATHWAY	mmu04510:Focal adhesion	0.021861 PGF, PDGFC, COL1A1, THBS4, SPP1	4.531047	0.949409
KEGG_PATHWAY	mmu04512:ECM-receptor interaction	0.076322 COL1A1, THBS4, SPP1	6.394956	0.999978
Enrichment Score:				
Annotation Cluster 12	1.2190157642888326	PValue Genes	Fold Enrich	Bonferroni
Category	Term			
UP_KEYWORDS	Zymogen	0.006696 CASP3, MMP9, CASP12, MMP3	10.14085	0.562359

GOTERM_MF_DIRECT	GO:0032403~protein complex binding	0.00971	CASP3, KRAS, MMP9, MMP3, APC	5.80141	0.718748
GOTERM_MF_DIRECT	GO:0004175~endopeptidase activity	0.028464	MMP9, CASP12, MMP3	11.12628	0.976577
UP_KEYWORDS	Protease	0.074212	CASP3, MMP9, CASP12, MMP3	3.98524	0.999924
	GO:0004252~serine-type				
GOTERM_MF_DIRECT	endopeptidase activity	0.088544	CASP3, MMP9, MMP3	5.878032	0.999994
GOTERM_MF_DIRECT	GO:0008233~peptidase activity	0.120434	CASP3, MMP9, CASP12, MMP3	3.220007	1
GOTERM_BP_DIRECT	GO:0006508~proteolysis	0.14461	CASP3, MMP9, CASP12, MMP3	2.958927	1
UP_KEYWORDS	Hydrolase	0.173852	PLA2G4A, CASP3, MMP9, ENDOG, CASP12, MMP3	1.968408	1
GOTERM_MF_DIRECT	GO:0016787~hydrolase activity	0.290242	PLA2G4A, CASP3, MMP9, ENDOG, CASP12, MMP3	1.625757	1

Enrichment Score:

Annotation Cluster 13 0.7639446711951424

Category	Term	PValue	Genes	Fold Enrich	Bonferroni
	GO:0051092~positive regulation of NF-				
GOTERM_BP_DIRECT	kappaB transcription factor activity	1.07E-04	KRAS, TNF, IL1RAP, IL1B, TLR4	19.7488	0.082689
UP_KEYWORDS	Innate immunity	0.009383	IL1RAP, TLR4, CD14, CSF1R	8.962656	0.686357
GOTERM_CC_DIRECT	GO:0009986~cell surface	0.009545	TNF, TLR4, PDGFC, CD14, IL1A, CSF1R	4.465592	0.661697
UP_KEYWORDS	Immunity	0.035645	IL1RAP, TLR4, CD14, CSF1R	5.386534	0.988488
GOTERM_BP_DIRECT	GO:0002376~immune system process	0.055627	IL1RAP, TLR4, CD14, CSF1R	4.496332	1
GOTERM_BP_DIRECT	GO:0045087~innate immune response	0.061758	IL1RAP, TLR4, CD14, CSF1R	4.305238	1
GOTERM_BP_DIRECT	GO:0007165~signal transduction	0.152515	KRAS, TNF, SOCS3, IL1RAP, TLR4, SMC3	2.058281	1
			TNF, TLR4, LIG4, HSP90B1, KRAS, ETS2, IL1RAP,		
GOTERM_CC_DIRECT	GO:0005886~plasma membrane	0.306227	PDGFC, IL1A, CD14, STAG2, CSF1R, APC	1.248637	1
UP_KEYWORDS	Cell membrane	0.530576	KRAS, TNF, IL1RAP, TLR4, PDGFC, CD14, CSF1R, APC	1.149242	1
			TNF, PGF, BNIP3, TLR4, FOS, HSP90B1, KRAS,		
GOTERM_CC_DIRECT	GO:0016020~membrane	0.749831	IL1RAP, PDGFC, STMN1, CD14, STAG2, CSF1R, APC	0.936553	1
UP_SEQ_FEATURE	topological domain:Extracellular	0.892681	TNF, IL1RAP, TLR4, CSF1R	0.778931	1
UP_KEYWORDS	Receptor	0.958207	IL1RAP, TLR4, CSF1R	0.619977	1
UP_SEQ_FEATURE	topological domain:Cytoplasmic	0.965466	TNF, IL1RAP, TLR4, CSF1R	0.610163	1
UP_SEQ_FEATURE	transmembrane region	0.993012	TNF, IL1RAP, BNIP3, TLR4, CSF1R	0.509412	1
			KRAS, TNF, IL1RAP, BNIP3, TLR4, PDGFC, CD14,		
UP_KEYWORDS	Membrane	0.997122	CSF1R, APC	0.559714	1
UP_KEYWORDS	Transmembrane helix	0.99966	TNF, IL1RAP, BNIP3, TLR4, CSF1R	0.389161	1

UP_KEYWORDS	Transmembrane GO:0016021~integral component of membrane	0.999671 TNF, IL1RAP, BNIP3, TLR4, CSF1R	0.38821	1
GOTERM_CC_DIRECT		0.999959 TNF, IL1RAP, BNIP3, TLR4, CSF1R	0.340319	1
Enrichment Score: Annotation Cluster 14 0.509790242861469				
Category Term PValue Genes Fold Enrich Bonferroni				
Category	Term	PValue	Genes	Fold Enrich Bonferroni
GOTERM_BP_DIRECT	GO:0007275~multicellular organism development	0.203126 TNF, PGF, PDGFC, STMN1, BMP5	2.091952	1
UP_KEYWORDS	Developmental protein	0.258344 PGF, PDGFC, STMN1, BMP5	2.213115	1
UP_KEYWORDS	Differentiation	0.326621 PGF, STMN1, BMP5	2.50774	1
GOTERM_BP_DIRECT	GO:0030154~cell differentiation	0.533127 PGF, STMN1, BMP5	1.655861	1
Enrichment Score: Annotation Cluster 15 0.48366828468521184				
Category Term PValue Genes Fold Enrich Bonferroni				
Category	Term	PValue	Genes	Fold Enrich Bonferroni
UP_KEYWORDS	Chromatin regulator	0.085546 KMT2C, PBRM1, SETD2	6	0.999983
GOTERM_BP_DIRECT	GO:0016569~covalent chromatin modification	0.121798 KMT2C, PBRM1, SETD2 CCNE2, FOS, NFKBIZ, ETS2, KMT2C, PBRM1, PDGFC,	4.855532	1
UP_KEYWORDS	Nucleus	0.181677 LIG4, SETD2, GADD45A, STAG2, SMC3	1.429202	1
UP_KEYWORDS	Activator	0.311894 NFKBIZ, KMT2C, SETD2	2.596154	1
UP_KEYWORDS	Transcription regulation	0.411349 NFKBIZ, ETS2, KMT2C, PBRM1, SETD2	1.500834	1
UP_KEYWORDS	Transcription	0.436409 NFKBIZ, ETS2, KMT2C, PBRM1, SETD2	1.452394	1
UP_KEYWORDS	DNA-binding	0.561786 FOS, ETS2, KMT2C, PBRM1	1.346633	1
GOTERM_BP_DIRECT	GO:0006355~regulation of transcription, DNA-templated	0.602023 FOS, NFKBIZ, ETS2, KMT2C, PBRM1, SETD2	1.133455	1
GOTERM_BP_DIRECT	GO:0006351~transcription, DNA-templated	0.631194 NFKBIZ, ETS2, KMT2C, PBRM1, SETD2	1.141973	1
GOTERM_MF_DIRECT	GO:0003677~DNA binding	0.643753 FOS, ETS2, KMT2C, PBRM1, LIG4	1.124475	1
Enrichment Score: Annotation Cluster 16 0.3683211269528496				
Category Term PValue Genes Fold Enrich Bonferroni				

GOTERM_CC_DIRECT	GO:0005925~focal adhesion binding site:ATP	0.195963 HSP90B1, KRAS, LIG4 0.373108 HSP90B1, LIG4, CSF1R	3.591889	1
UP_SEQ_FEATURE	Nucleotide-binding	0.392446 HSP90B1, KRAS, LIG4, SMC3, CSF1R	2.260637	1
UP_KEYWORDS	ATP-binding	0.450708 HSP90B1, LIG4, SMC3, CSF1R	1.539339	1
GOTERM_MF_DIRECT	GO:0000166~nucleotide binding	0.681485 HSP90B1, KRAS, LIG4, SMC3, CSF1R	1.072781	1
GOTERM_MF_DIRECT	GO:0005524~ATP binding	0.699734 HSP90B1, LIG4, SMC3, CSF1R	1.102537	1

Enrichment Score:

Annotation Cluster 17 0.1231810958443257

Category	Term	PValue	Genes	Fold Enrich	Bonferroni
UP_KEYWORDS	Metal-binding	0.590885	PLA2G4A, MMP9, ENDOG, KMT2C, COL1A1, LIG4, MMP3	1.113402	1
GOTERM_MF_DIRECT	GO:0008270~zinc ion binding	0.728182	MMP9, KMT2C, MMP3	1.159203	1
GOTERM_MF_DIRECT	GO:0046872~metal ion binding	0.827141	PLA2G4A, MMP9, ENDOG, KMT2C, COL1A1, LIG4, MMP3	0.866667	1
UP_KEYWORDS	Zinc	0.903552	MMP9, KMT2C, MMP3	0.771796	1