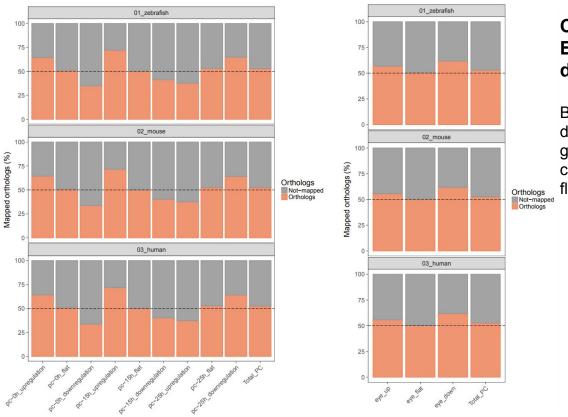
Orthologs

Number of orthologs in the different gene sets

Our regeneration Data:

High conservation of upregulated genes compared to the other gene sets in the early time points



Comparison: Eye development

Both up and down-regulated genes are more conserved than flat genes

Chi square test

Our regeneration data

	Human		Mouse		Zebrafish	
	up	down	up	down	up	down
0h	0	0,00001807	0	0,00002067	0	0,00016437
15h	0	0,00610516	0	0,00263558	0	0,01103296
25h	0	1,70E-07	0	1,40E-07	0	6,00E-08

	Human	Mouse			Zebrafish	
	up	down	up	down	up	down
0h	122	18	130	18	123	14,2
15h	271	8	243	9	272	6,46
25h	57	27	54	28	56,46	29

Comparison: Eye development

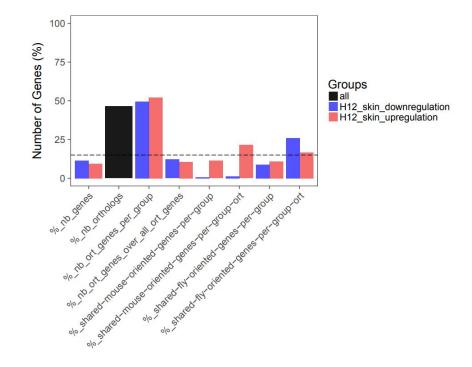
	Human		Mouse		Zebrafish	
	up	down	up	down	up	down
chi-test	17	83	17	83	23,76	79
p-value	0,00003467	0	0,00003561	0	0,00000109	0

Baseline: flat genes

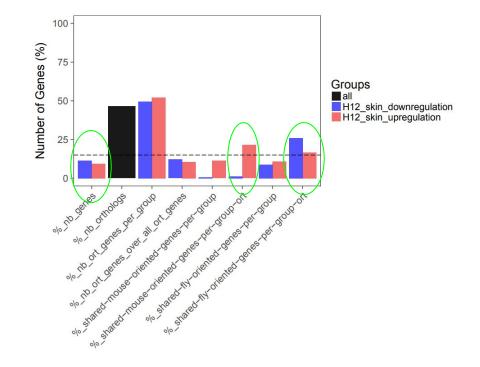
Goal comparison to eye: we would like to find a negative control, however the eye development is conserved [1]. We were not able to find a negative control.

[1] <u>http://dx.doi.org/10.1016/S0960-9822(00)00821-6</u>

Mouse skin regeneration data



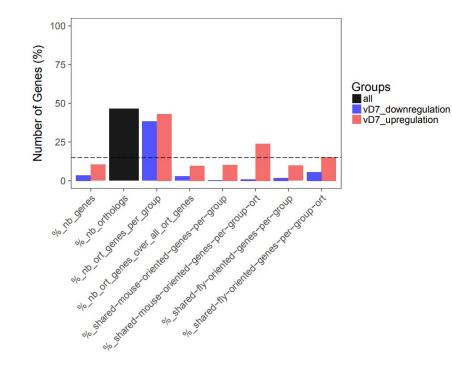
Mouse skin regeneration data



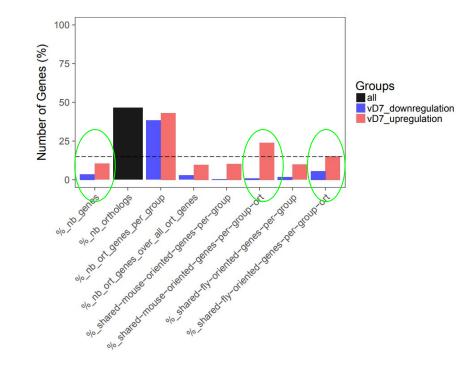
Mouse skin

Category	Term	Count	PValue
OTERM_CC_DIRECT	GO:0005737~cytoplasm	51	1,12E+11
OTERM_BP_DIRECT	GO:0006955~immune response	8	4,06E+10
OTERM_MF_DIRECT	GO:0005515~protein binding	27	1,85E+11
OTERM_MF_DIRECT	GO:0016019~peptidoglycan receptor activity	3	0.0010467788627076806
OTERM_BP_DIRECT	GO:0001751~compound eye photoreceptor cell differentiation	5	0.001354152208753223
OTERM_CC_DIRECT	GO:0016020~membrane	17	0.001462933721966101
OTERM_BP_DIRECT	GO:0007476~imaginal disc-derived wing morphogenesis	12	0.00154206663154347
OTERM_MF_DIRECT	GO:0050839~cell adhesion molecule binding	5	0.00200454137001363
OTERM_MF_DIRECT	GO:0008745~N-acetylmuramoyl-L-alanine amidase activity	4	0.002056336522177214
OTERM_BP_DIRECT	GO:0009253~peptidoglycan catabolic process	4	0.002073126267653188
OTERM_BP_DIRECT	GO:0045087~innate immune response	8	0.002330633090130692
EGG_PATHWAY	dme04144:Endocytosis	10	0.002617007083932810
OTERM_BP_DIRECT	GO:0008063~Toll signaling pathway	5	0.003147341481316913
OTERM_MF_DIRECT	GO:0042834~peptidoglycan binding	4	0.004361492653905138
OTERM_CC_DIRECT	GO:0031594~neuromuscular junction	6	0.004444336794147326
OTERM_BP_DIRECT	GO:0007298~border follicle cell migration	8	0.008257629762045353
OTERM_CC_DIRECT	GO:0005925~focal adhesion	4	0.00846532154636451
OTERM_BP_DIRECT	GO:0007616~long-term memory	6	0.00856107336624281
OTERM CC DIRECT	GO:0005912~adherens junction	5	0.011720489804138198
OTERM_BP_DIRECT	GO:0009950~dorsal/ventral axis specification	4	0.012629891923736493
OTERM_BP_DIRECT	GO:0055085~transmembrane transport	12	0.013408701411461315
OTERM_BP_DIRECT	GO:0008299~isoprenoid biosynthetic process	3	0.015188480189017278
OTERM BP DIRECT	GO:0006950~response to stress	3	0.015188480189017278
OTERM_BP_DIRECT	GO:0006964~positive regulation of biosynthetic process of antibacterial peptides activ	3	0.018017762245826307
OTERM_CC_DIRECT	GO:0000151~ubiquitin ligase complex	4	0.01870968431927909
OTERM_BP_DIRECT	GO:0035220~wing disc development	6	0.01917905130958334
OTERM BP DIRECT	GO:0030431~sleep	7	0.020436893387287748
OTERM_CC_DIRECT	GO:0005886~plasma membrane	23	0.02185287794818437
OTERM BP DIRECT	GO:0007465~R7 cell fate commitment	3	0.024278997219393254
OTERM BP DIRECT	GO:0007155~cell adhesion	6	0.02472742152681488
	GO:0070374~positive regulation of ERK1 and ERK2 cascade	4	0.024860952835202482
	GO:0005887~integral component of plasma membrane	16	0.02780013640225199
	GO:0007423~sensory organ development	5	0.029918315731241547
OTERM BP DIRECT	GO:0007464~R3/R4 cell fate commitment	3	0.031291927223918325
	GO:0010025~wax biosynthetic process	3	0.031291927223918325
	GO:0035195~gene silencing by miRNA	3	0.031291927223918325
	GO:0035336~long-chain fatty-acyl-CoA metabolic process	3	0.031291927223918325
	GO:0043195~terminal bouton	4	0.03363727855494449
	GO:0002781~antifungal peptide production	2	0.034797175408941665
	GO:0000184~nuclear-transcribed mRNA catabolic process, nonsense-mediated deca	3	0.03506161002368078
	GO:0046579~positive regulation of Ras protein signal transduction	4	0.03705084443634689
	GO:0052692~raffinose alpha-galactosidase activity	2	0.037345951142604576
		-	0.001010001112004010
	GO:0004557~alpha-galactosidase activity	2	0.037345951142604576

Mouse heart regeneration data



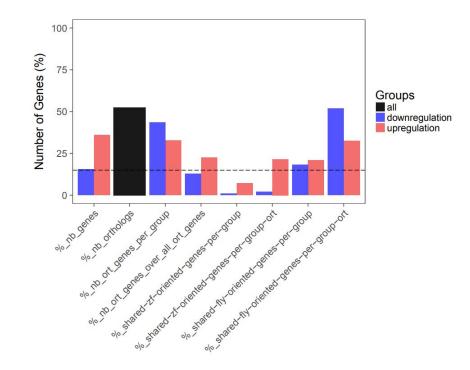
Mouse heart regeneration data



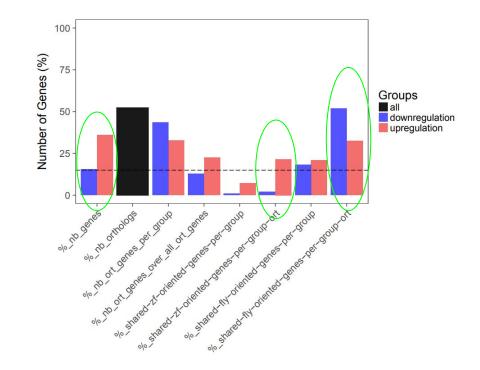
Mouse heart

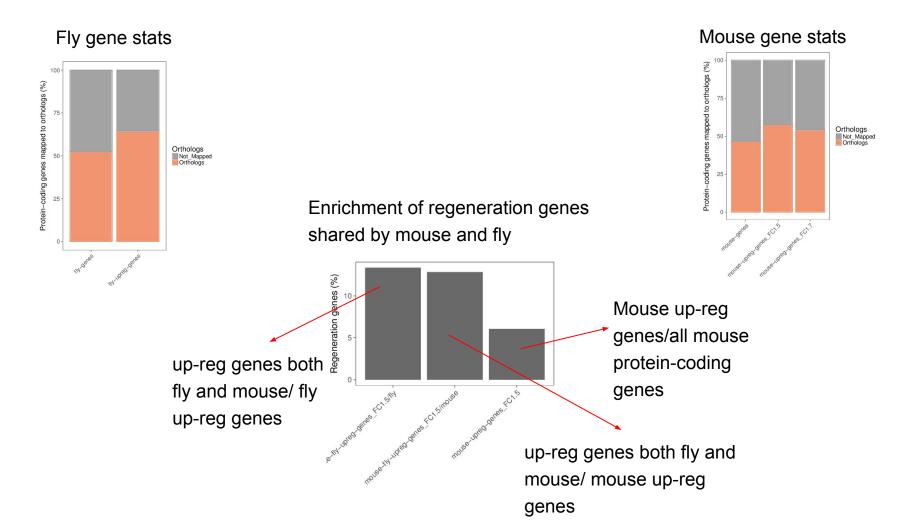
Category	Term	PValue
GOTERM_BP_DIRECT	GO:0007422~peripheral nervous system development	5,86E+08
GOTERM_BP_DIRECT	GO:0007307~eggshell chorion gene amplification	7,90E+08
KEGG_PATHWAY	dme00830:Retinol metabolism	1,20E+12
GOTERM_BP_DIRECT	GO:0009253~peptidoglycan catabolic process	2.64E+11
GOTERM_MF_DIRECT	GO:0016019~peptidoglycan receptor activity	2,71E+12
GOTERM_MF_DIRECT	GO:0008745~N-acetylmuramoyl-L-alanine amidase activity	2.87E+11
KEGG_PATHWAY	dme00983:Drug metabolism - other enzymes	6.18E+11
GOTERM_MF_DIRECT	GO:0042834~peptidoglycan binding	6.26E+11
GOTERM_MF_DIRECT	GO:0016758~transferase activity, transferring hexosyl groups	6.69E+11
GOTERM_MF_DIRECT	GO:0005515~protein binding	0.0011433108656887288
GOTERM_BP_DIRECT	GO:0007155~cell adhesion	
GOTERM_BP_DIRECT	GO:0009813~flavonoid biosynthetic process	0.001296374949553805
GOTERM_BP_DIRECT	GO:0052696~flavonoid glucuronidation	0.003079095812116281
GOTERM_BP_DIRECT	GO:0007095~mitotic G2 DNA damage checkpoint	0.003079095812116281
GOTERM_BP_DIRECT	GO:0008284~positive regulation of cell proliferation	0.0034348830497672738
GOTERM_BP_DIRECT	GO:0000082~G1/S transition of mitotic cell cycle	0.003924133701479878
GOTERM_MF_DIRECT	GO:0008194~UDP-glycosyltransferase activity	0.004607609546810708
GOTERM_BP_DIRECT	GO:0042127~regulation of cell proliferation	0.004838406242117356
GOTERM_BP_DIRECT	GO:0007419~ventral cord development	0.005620441227803466
GOTERM_MF_DIRECT	GO:0003677~DNA binding	0.005956109539193618
GOTERM_MF_DIRECT	GO:0008028~monocarboxylic acid transmembrane transporter activity	0.006452392487151536
GOTERM_MF_DIRECT	GO:0016538~cyclin-dependent protein serine/threonine kinase regulator activity	0.006617673779342341
GOTERM_MF_DIRECT	GO:0003678~DNA helicase activity	0.006617673779342341
GOTERM_BP_DIRECT	GO:0022008~neurogenesis	0.007672753657145244
GOTERM_BP_DIRECT	GO:0006260~DNA replication	0.009301236943766104
GOTERM_CC_DIRECT	GO:0005667~transcription factor complex	0.009642488227042278
GOTERM_BP_DIRECT	GO:0007476~imaginal disc-derived wing morphogenesis	0.010422904774709814
GOTERM_BP_DIRECT	GO:0006955~immune response	0.011171192066803972
GOTERM_BP_DIRECT	GO:0045087~innate immune response	0.011273788270270256
GOTERM_MF_DIRECT	GO:0001077~transcriptional activator activity, RNA polymerase II core promoter proximal re	
GOTERM_BP_DIRECT	GO:0000915~actomyosin contractile ring assembly	0.012460793699813105
GOTERM_MF_DIRECT	GO:0003700~transcription factor activity, sequence-specific DNA binding	0.013419058710738313
GOTERM_BP_DIRECT	GO:0010971~positive regulation of G2/M transition of mitotic cell cycle	0.013939495616176752
GOTERM_BP_DIRECT	GO:1900117~regulation of execution phase of apoptosis	0.016190497508208754
GOTERM_MF_DIRECT	GO:0016614~oxidoreductase activity, acting on CH-OH group of donors	0.017205148476260687
GOTERM_BP_DIRECT	GO:000281~mitotic cytokinesis	0.017205148476260687
GOTERM_BP_DIRECT	GO:0048477~oogenesis	0.018534707050843684
GOTERM_BP_DIRECT	GO:0043066~negative regulation of apoptotic process	0.018551290314511068
GOTERM_CC_DIRECT	GO:0005634~nucleus	0.01911653879914678
GOTERM_BP_DIRECT	GO:0002009~morphogenesis of an epithelium	0.01931550508090287
GOTERM_BP_DIRECT	GO:0007400~neuroblast fate determination	0.01954001102585428
GOTERM_CC_DIRECT	GO:0043231~intracellular membrane-bounded organelle	0.020979684425786856
GOTERM_BP_DIRECT	GO:0035561~regulation of chromatin binding	0.020979684425786856

Zebrafish heart regeneration data

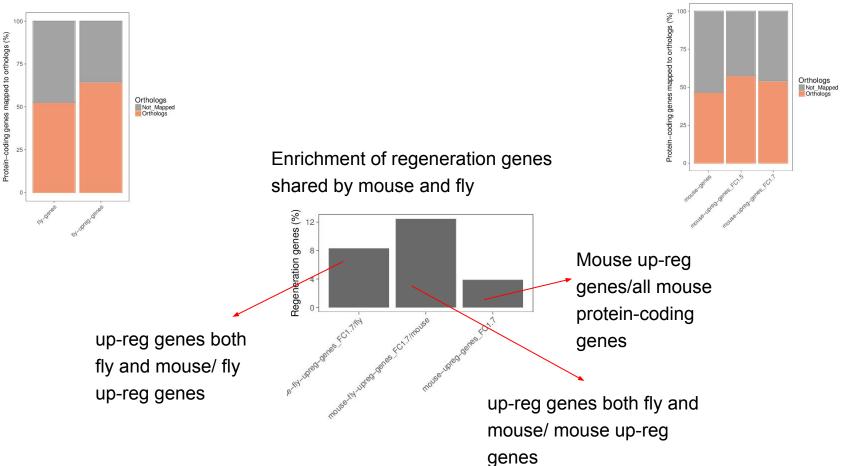


Zebrafish heart regeneration data





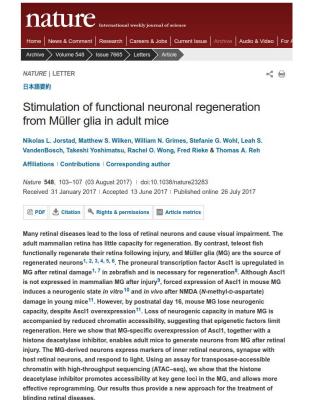
Fly gene stats

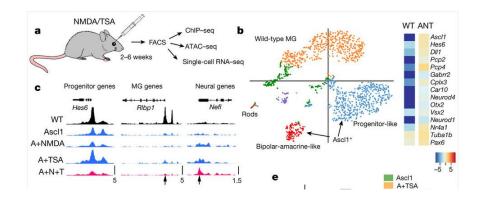


Mouse gene stats

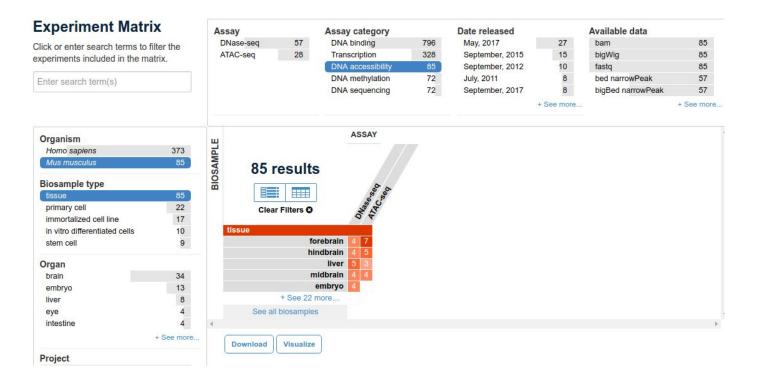
New data for re-usage analysis

New data for re-usage analysis





ENCODE data for re-usage analysis



ENCODE data for re-usage analysis

ATAC-seq of midbrain	Experiment
Mus musculus C57BL/6 midbrain embryo (16.5 days) Lab: Bing Ren, UCSD Project: ENCODE	ENCSR096JCC released
ATAC-seq of midbrain	Experiment
Mus musculus C57BL/6 midbrain embryo (15.5 days) Lab: Bing Ren, UCSD Project: ENCODE	ENCSR468GUI released
ATAC-seq of midbrain	Experiment
Mus musculus C57BL/6 midbrain embryo (13.5 days) Lab: Bing Ren, UCSD Project: ENCODE	ENCSR819QOJ released
ATAC-seq of midbrain	Experiment
Mus musculus C57BL/6 midbrain embryo (11.5 days)	ENCSR382RUC
Lab: Bing Ren, UCSD Project: ENCODE	released

ATAC-seq of liver	Experiment
Mus musculus C57BL/6 liver postnatal (0 days)	ENCSR609OHJ
Lab: Bing Ren, UCSD	released
Project: ENCODE	
ATAC-seq of liver	Experiment
Mus musculus C57BL/6 liver embryo (12.5 days)	ENCSR302LIV
Lab: Bing Ren, UCSD	released
Project: ENCODE	
ATAC-seq of liver	Experiment
Mus musculus C57BL/6 liver embryo (11.5 days)	ENCSR785NEL
Lab: Bing Ren, UCSD	released
Project: ENCODE	

Genes in/out side clusters

Oh upregulated genes INSIDE clusters

Category	Term	Count	PValue
GOTERM_BP_DIRECT	GO:0030707~ovarian follicle cell development	17	2,21E+08
GOTERM_BP_DIRECT	GO:0007399~nervous system development	18	7,96E+08
GOTERM_MF_DIRECT	GO:0003677~DNA binding	45	8,76E+08
GOTERM_BP_DIRECT	GO:0007095~mitotic G2 DNA damage checkpoint	13	1,15E+11
GOTERM_BP_DIRECT	GO:0048477~oogenesis	27	1,10E+12
GOTERM_BP_DIRECT	GO:0046425~regulation of JAK-STAT cascade	6	2,11E+12
GOTERM_CC_DIRECT	GO:0005634~nucleus	100	2,78E+12
GOTERM_BP_DIRECT	GO:0007219~Notch signaling pathway	11	2,97E+11
GOTERM_CC_DIRECT	GO:0005737~cytoplasm	87	3,54E+11
GOTERM_BP_DIRECT	GO:0035019~somatic stem cell population maintenance	6	5,13E+11
GOTERM_BP_DIRECT	GO:0051123~RNA polymerase II transcriptional preinitiation complex	6	8,44E+11
GOTERM_MF_DIRECT	GO:0004602~glutathione peroxidase activity	7	0.0010275094378868212
GOTERM_CC_DIRECT	GO:0005700~polytene chromosome	17	0.001825478636670938
GOTERM_BP_DIRECT	GO:0007423~sensory organ development	9	0.0031706855101392814
GOTERM_BP_DIRECT	GO:0008283~cell proliferation	9	0.0031706855101392814
GOTERM_BP_DIRECT	GO:0000278~mitotic cell cycle	8	0.0039569142473899165
GOTERM_BP_DIRECT	GO:0007365~periodic partitioning	3	0.003999565424569938
GOTERM_MF_DIRECT	GO:0003682~chromatin binding	13	0.005391939827190603
GOTERM_BP_DIRECT	GO:0007067~mitotic nuclear division	13	0.005413548534288489
GOTERM_BP_DIRECT	GO:0019991~septate junction assembly	6	0.005980092676411749
GOTERM_BP_DIRECT	GO:0001708~cell fate specification	6	0.005980092676411749
GOTERM_MF_DIRECT	GO:0005515~protein binding	40	0.006229503724949665
GOTERM_BP_DIRECT	GO:0044804~nucleophagy	4	0.0066527383680596245
GOTERM_BP_DIRECT	GO:0042058~regulation of epidermal growth factor receptor signaling	4	0.0066527383680596245
GOTERM_MF_DIRECT	GO:0003700~transcription factor activity, sequence-specific DNA bind	21	0.0071006277023068845
GOTERM_BP_DIRECT	GO:0006749~glutathione metabolic process	7	0.007367015961335778
GOTERM_MF_DIRECT	GO:0016301~kinase activity	7	0.007637760324176527
GOTERM_BP_DIRECT	GO:0000706~meiotic DNA double-strand break processing	3	0.007803286413356147
GOTERM_BP_DIRECT	GO:0006367~transcription initiation from RNA polymerase II promoter	8	0.009126199902704371
GOTERM_MF_DIRECT	GO:0000978~RNA polymerase II core promoter proximal region sequ	5	0.010364860296567718
GOTERM_BP_DIRECT	GO:0006357~regulation of transcription from RNA polymerase II pron	14	0.012033098157289333
GOTERM_BP_DIRECT	GO:0006351~transcription, DNA-templated	24	0.012497319333413156
GOTERM_BP_DIRECT	GO:0042059~negative regulation of epidermal growth factor receptor	6	0.01385007273141317
GOTERM_CC_DIRECT	GO:0019008~molybdopterin synthase complex	3	0.014096912478004651
GOTERM_MF_DIRECT	GO:0001046~core promoter sequence-specific DNA binding	3	0.014698540863274719
GOTERM_MF_DIRECT	GO:0001077~transcriptional activator activity, RNA polymerase II con	7	0.016886167182106705
GOTERM_BP_DIRECT	GO:0006777~Mo-molybdopterin cofactor biosynthetic process	3	0.01856901108354489
GOTERM_BP_DIRECT	GO:0046579~positive regulation of Ras protein signal transduction	6	0.018705690772873797
GOTERM_BP_DIRECT	GO:0006468~protein phosphorylation	18	0.02010765351329353
GOTERM_MF_DIRECT	GO:0046872~metal ion binding	38	0.020489186510212913
KEGG_PATHWAY	dme00980:Metabolism of xenobiotics by cytochrome P450	8	0.02063220607153842

Oh upregulated genes OUTSIDE clusters

Category	Term	Count	PValue
GOTERM_MF_DIRECT	GO:0005515~protein binding	119	4,57E+06
GOTERM_BP_DIRECT	GO:0016567~protein ubiquitination	39	3,90E+07
GOTERM_MF_DIRECT	GO:0004842~ubiquitin-protein transferase activity	44	3,47E+08
GOTERM_CC_DIRECT	GO:0005737~cytoplasm	221	7,69E+09
GOTERM_BP_DIRECT	GO:0000122~negative regulation of transcription from RNA polymerase II prom	33	3,57E+10
GOTERM_BP_DIRECT	GO:0007411~axon guidance	38	4,73E+10
GOTERM_BP_DIRECT	GO:0007476~imaginal disc-derived wing morphogenesis	42	7,39E+10
GOTERM_CC_DIRECT	GO:0005634~nucleus	248	7,58E+10
GOTERM_MF_DIRECT	GO:0008270~zinc ion binding	100	1,76E+12
GOTERM_MF_DIRECT	GO:0031624~ubiquitin conjugating enzyme binding	9	1,88E+12
GOTERM_CC_DIRECT	GO:0000151~ubiquitin ligase complex	12	3,63E+11
GOTERM_BP_DIRECT	GO:0001738~morphogenesis of a polarized epithelium	8	5,73E+11
GOTERM_BP_DIRECT	GO:0045892~negative regulation of transcription, DNA-templated	25	9,65E+11
GOTERM_MF_DIRECT	GO:0061630~ubiquitin protein ligase activity	19	0.001064090530109791
GOTERM_BP_DIRECT	GO:0000209~protein polyubiquitination	16	0.001171373940860710
GOTERM_BP_DIRECT	GO:0006897~endocytosis	16	0.001394874217457518
GOTERM_BP_DIRECT	GO:0007391~dorsal closure	22	0.001517361706241614
GOTERM BP_DIRECT	GO:0045167~asymmetric protein localization involved in cell fate determination	7	0.001647727692461506
GOTERM BP_DIRECT	GO:0042787~protein ubiquitination involved in ubiquitin-dependent protein cata	19	0.001792036409709696
GOTERM BP DIRECT	GO:0035050~embryonic heart tube development	8	0.00191837279806341
GOTERM CC DIRECT	GO:0032154~cleavage furrow	11	0.002008641954260344
GOTERM_BP_DIRECT	GO:0043066~negative regulation of apoptotic process	16	0.002289973959461549
GOTERM BP DIRECT	GO:0006909~phagocytosis	36	0.002293781737628386
GOTERM MF DIRECT	GO:0005524~ATP binding	107	0.002372486021273864
GOTERM_BP_DIRECT	GO:0045746~negative regulation of Notch signaling pathway	13	0.002640038757936424
GOTERM BP DIRECT	GO:0006727~ommochrome biosynthetic process	8	0.002693731617038300
GOTERM BP DIRECT	GO:0030837~negative regulation of actin filament polymerization	5	0.002959338070500696
GOTERM_BP_DIRECT	GO:1902669~positive regulation of axon guidance	5	0.002959338070500696
GOTERM_BP_DIRECT	GO:0022008~neurogenesis	79	0.003409904063745352
KEGG_PATHWAY	dme00563:Glycosylphosphatidylinositol(GPI)-anchor biosynthesis	8	0.003510628849266861
GOTERM_BP_DIRECT	GO:0051301~cell division	16	0.003616217855481752
GOTERM BP DIRECT	GO:0007307~equipments and a second se	8	0.003688562838048898
GOTERM_CC_DIRECT	GO:0005794~Golgi apparatus	29	0.004431059100334946
GOTERM_BP_DIRECT	GO:0007422~peripheral nervous system development	18	0 004441844061441030
GOTERM BP DIRECT	GO:0008360~regulation of cell shape	21	0.004446220157697897
GOTERM BP DIRECT	GO:000082~G1/S transition of mitotic cell cvcle	6	0.004756767312969577
GOTERM BP DIRECT	GO:0006886~intracellular protein transport	21	0.004967734767808359
GOTERM MF DIRECT	GO:0008134~transcription factor binding	19	0.00517994844073751
GOTERM_BP_DIRECT	GO:0007029~endoplasmic reticulum organization	7	0.005190886947870575
GOTERM_BP_DIRECT	GO:0045773~positive regulation of axon extension	5	0.005441147136539997
GOTERM CC DIRECT	GO:0045179~apical cortex	11	0.006199954132911566
JOILAM_CO_DIRECT	CO. CONTRA aproal CONTEX		0.000100004102011000

Orthologous genes in 0h upregulated clusters

