

Supplementary Materials for

UVB Irradiation–Induced Pain Is Mediated by CXCL5

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Fig. S1. Validation of up-regulated transcripts (as measured by PCR array cards) with qPCR.

Fig. S2. Correlation of PCR array data between male and female volunteers.

Fig. S3. The increased expression of CXCL5 mRNA is attenuated in the UVB model by piroxicam treatment.

Fig. S4. UVB-induced changes in female rats.

Table S1. Rat PCR array data.

Table S2. Human PCR array data.

Supplementary Material

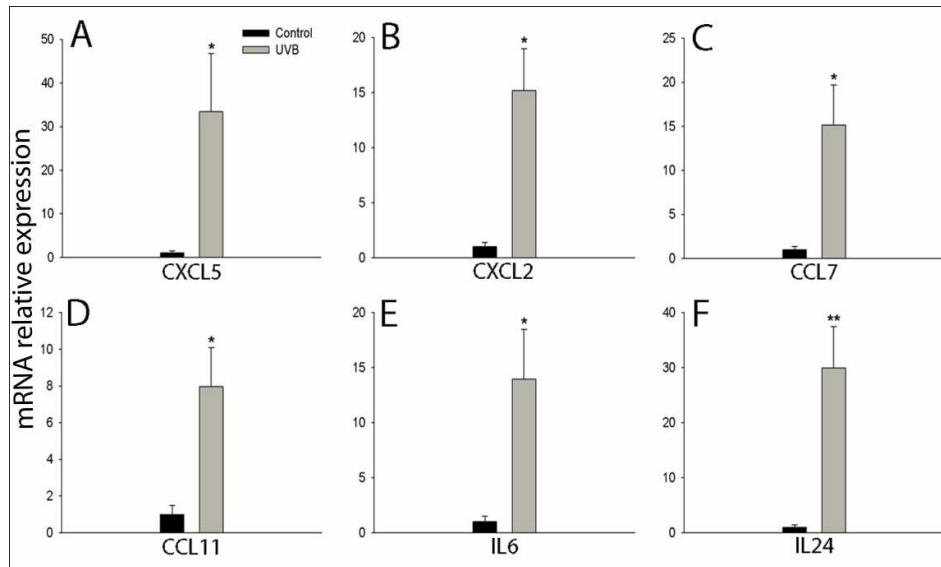


Fig. S1. Validation of up-regulated transcripts (as measured by PCR array cards) with qPCR. A significant increase in transcript levels was measured for CXCL5, CXCL2, CCL7, CCL11, IL6 and IL24 in UVB treated versus non-irradiated contralateral rat skin (A-F, respectively). Wilcoxon signed rank test (A), paired t-test (B-F), n=7. Mean \pm SEM, *p<0.05 **p<0.01.

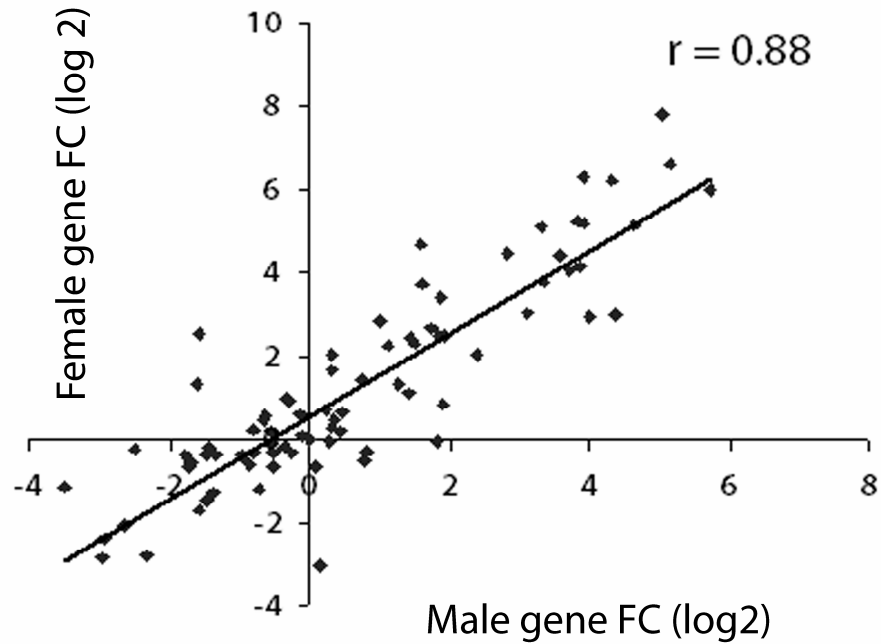


Fig. S2. Correlation of PCR array data between male and female volunteers.

Gene expression after UVB irradiation for male human subjects was plotted against the data from females. The Pearson's correlation coefficient, $r = 0.88$, showed that there is a strong positive relationship between the two data sets. This relationship is also significant with $p < 0.001$. (Males $n=5$, Female $n=3$, genes $n=92$).

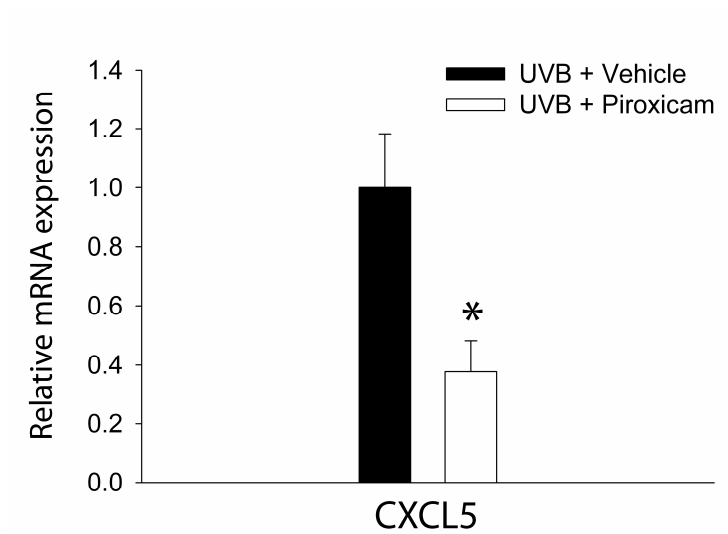


Fig. S3. The increased expression of CXCL5 mRNA is attenuated in the UVB model by piroxicam treatment. Just prior to irradiation and at 12, 24, 36 and 46 hours after UVB treatment, rats received an i.p. injection of either Piroxicam (Sigma, 10mg/kg) or vehicle (20% DMSO). Rats treated with Piroxicam had significantly reduced CXCL5 expression in UVB treated skin compared those in the vehicle group. (* $p < 0.05$, t-test, $n = 4$). Data are presented as mean \pm SEM.

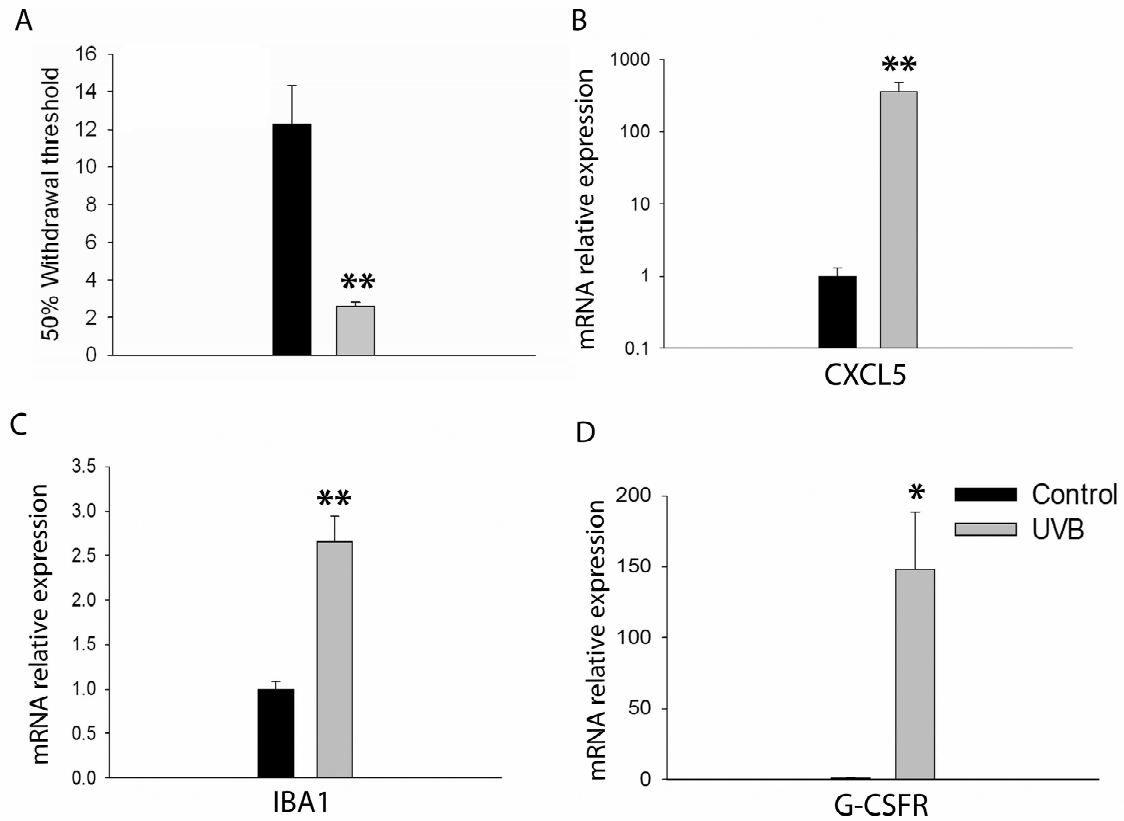


Fig. S4. UVB-induced changes in female rats. (A) Female rats developed a significant mechanical pain-related hypersensitivity 48 hours after UVB treatment. UVB treatment also significantly increased the mRNA expression of CXCL5 (B), IBA1 (C) and GCSFR (D). Paired t-test, * $p < 0.05$, ** $p < 0.01$, paired t-test $n = 6$. All data are presented as the mean \pm SEM.

Table S1. Rat PCR array data.

Applied Biosystem primer code; Rat	Name	FC	
		FINAL	adjPVal
Alox15-Rn00696151_m1	Arachidonate 15-Lipoxygenase	6.88504	0.121519
Alox5-Rn00563172_m1	Arachidonate 5-lipoxygenase	2.117766	8.37E-05
Areg-Rn00567471_m1	Amphiregulin	0.90579	0.948159
Artn-Rn01761472_g1	Artemin	1.686631	0.035249
Bdnf-Rn01484928_m1	Brain-derived neurotrophic factor	2.47553	0.325985
Btc-Rn00673281_m1	Betacellulin	0.369867	0.338399
C3-Rn00566466_m1	Complement component 3	4.971303	0.000131
C5-Rn01436156_m1	Complement component 5	1.93606	0.443386
Ccl1-Rn01752376_m1	Chemokine (C-C motif) ligand 1	0.354474	0.017597
Ccl11-Rn00569995_m1	Chemokine (C-C motif) ligand 11	11.58508	4.51E-05
Ccl17-Rn00589290_m1	Chemokine (C-C motif) ligand 17	1.142472	0.586833
Ccl19-Rn01439563_m1	Chemokine (C-C motif) ligand 19	2.755498	0.016383
Ccl2-Rn00580555_m1	Chemokine (C-C motif) ligand 2	14.66058	0.000221
Ccl20-Rn00570287_m1	Chemokine (C-C motif) ligand 20	0.353528	0.058643
Ccl21b-Rn01764651_g1	Chemokine (C-C motif) ligand 21b	0.755235	0.291932
Ccl22-Rn01536591_m1	Chemokine (C-C motif) ligand 22	0.137293	1.14E-05
Ccl24-Rn01481451_m1	Chemokine (C-C motif) ligand 24	1.344512	0.187421
Ccl25-Rn01403351_m1	Chemokine (C-C motif) ligand 25	0.57458	0.44041

Ccl26-Rn01481484_m1	Chemokine (C-C motif) ligand 26	NA	NA
Ccl27-Rn01437037_m1	Chemokine (C-C motif) ligand 27	NA	NA
Ccl28-Rn00586715_m1	Chemokine (C-C motif) ligand 28	NA	NA
Ccl3-Rn00564660_m1	Chemokine (C-C motif) ligand 3	5.997417	0.007584
Ccl4-Rn00587826_m1	Chemokine (C-C motif) ligand 4	15.35799	0.000137
Ccl5-Rn00579590_m1	Chemokine (C-C motif) ligand 5	2.664209	0.099169
Ccl6-Rn01456402_g1	Chemokine (C-C motif) ligand 6	2.455923	0.004211
Ccl7-Rn01467286_m1	Chemokine (C-C motif) ligand 7	14.232	0.000131
Ccl9-Rn01471276_m1	Chemokine (C-C motif) ligand 9	6.526384	0.000137
Csf1-Rn00576849_m1	Colony stimulating factor 1 (macrophage)	4.542358	1.14E-05
Csf2-Rn01456851_m1	Colony stimulating factor 2 (granulocyte-macrophage)	0.87465	0.918736
Csf3-Rn00567344_m1	Colony stimulating factor 3 (granulocyte)	7.089967	0.092509
Cx3cl1-Rn00593186_m1	Chemokine (C-X3-C motif) ligand 1	0.472526	0.001883
Cxcl1-Rn00578225_m1	Chemokine (C-X-C motif) ligand 1	5.410761	0.006414
Cxcl10-Rn00594648_m1	Chemokine (C-X-C motif) ligand 10	2.225852	0.271036
Cxcl11-Rn00788262_g1	Chemokine (C-X-C motif) ligand 11	3.133687	0.19442
Cxcl12-Rn00573260_m1	Chemokine (C-X-C motif) ligand 12	1.576372	0.03271
Cxcl13-Rn01450028_m1	Chemokine (C-X-C motif) ligand 13	3.053277	0.012622
Cxcl14-Rn01441840_m1	Chemokine (C-X-C motif) ligand 14	1.885975	0.003849
Cxcl16-Rn01496393_m1	Chemokine (C-X-C motif) ligand 16	1.221816	0.34029
Cxcl17-Rn01764053_m1	Chemokine (C-X-C motif) ligand 17	0.95014	0.958139
Cxcl2-Rn00586403_m1	Chemokine (C-X-C motif) ligand 2	24.6661	0.009162
Cxcl3-Rn00593435_m1	Chemokine (C-X-C motif) ligand 3	2.05144	0.294366
Cxcl5-Rn00573587_g1	Chemokine (C-X-C motif) ligand 5	51.28608	0.000171
Cxcl9-Rn00595504_m1	Chemokine (C-X-C motif) ligand 9	1.0446	0.897726
Ebi3-Rn01527778_m1	Epstein-Barr virus induced gene 3	4.03549	0.000469
Edn1-Rn00561129_m1	Endothelin 1	1.07337	0.897726
Ereg-Rn00572454_m1	Epiregulin	0.89458	0.918963
Fgf7-Rn00573319_m1	Fibroblast growth factor 7 (keratinocyte growth factor)	5.80183	0.000183
Hbegf-Rn00564075_m1	Heparin-binding EGF-like growth factor	4.28442	0.009229
Ifn-g-Rn00594078_m1	Interferon, gamma	0.58442	0.437389
Il10-Rn01644839_m1	Interleukin 10	10.6731	0.00088
Il11-Rn00591721_m1	Interleukin 11	0.66565	0.106098
Il12a-Rn00584538_m1	Interleukin 12a	0.536757	0.002136
Il12b-Rn00575112_m1	Interleukin 12b	0.277929	0.037099

Il13-Rn00587615_m1	Interleukin 13	1.01954	0.96705
Il15-Rn00689964_m1	Interleukin 15	1.03152	0.946766
Il16-Rn01477722_m1	Interleukin 16	0.80001	0.293906
Il18-Rn01422083_m1	Interleukin 18	0.63103	0.184776
Il19-Rn01490483_m1	Interleukin 19	6.17235	0.002319
Il1a-Rn00566700_m1	Interleukin 1 alpha	0.2102	0.000155
Il1b-Rn00580432_m1	Interleukin 1 beta	4.9379	0.026934
Il2-Rn99999181_m1	Interleukin 2	NA	NA
Il20-Rn01483978_m1	Interleukin 20	0.05706	1.72E-05
Il21-Rn01755623_m1	Interleukin 21	1.36465	0.506481
Il23a-Rn00590334_g1	Interleukin 23a	1.48595	0.26231
Il24-Rn00591159_m1	Interleukin 24	32.69474	0.000417
Il27-Rn01510484_m1	Interleukin 27	1.075036	0.928099
Il3-Rn00580435_m1	Interleukin 3	8.93579	0.001386
Il33-Rn01759837_m1	Interleukin 33	2.398043	0.000198
Il4-Rn01456866_m1	Interleukin 4	NA	NA
Il5-Rn01459975_m1	Interleukin 5	NA	NA
Il6-Rn00561420_m1	Interleukin 6	14.80302	0.001406
Il7-Rn00681900_m1	Interleukin 7	1.297176	0.291932
Il9-Rn01448718_m1	Interleukin 9	NA	NA
Lif-Rn00573491_g1	Leukemia inhibitory factor	21.35104	0.019041
LOC301289-Rn01757168_m1	Similar to Interleukin-17 precursor (IL-17) (Cytotoxic T lymphocyte-associated antigen 8) (CTLA-8)	NA	NA
LOC682457-Rn01516136_m1	Taxilin alpha (interleukin 14)	0.866346	0.410661
Lta4h-Rn01503878_m1	Leukotriene A4 hydrolase	0.980934	0.928099
Mif-Rn00821234_g1	Macrophage migration inhibitory factor	1.125275	0.291932
Ngf-Rn01533872_m1	Nerve growth factor (beta polypeptide)	1.41969	0.198758
Nos2-Rn00561646_m1	Nitric oxide synthase 2, inducible	34.29193	0.010603
Pf4-Rn01768298_g1	Platelet factor 4 (CXCL4)	5.986383	1.72E-05
Ppbp-Rn00596603_g1	Pro-platelet basic protein (chemokine (C-X-C motif) ligand 7)	14.02202	0.007272
Ptges-Rn00572047_m1	Prostaglandin E synthase	1.057634	0.918736
Ptgis-Rn00580687_m1	Prostaglandin I2 (prostacyclin) synthase	4.254087	0.000198
Ptgs2-Rn01483828_m1	Prostaglandin-endoperoxide synthase 2 (cyclooxygenase-2)	4.60403	0.00309
Tgfb1-Rn00572010_m1	Transforming growth factor, beta 1	1.792186	0.014843

Tnf-Rn99999017_m1	Tumor necrosis factor	1.52635	0.191037
Xcl1-Rn00592605_m1	Chemokine (C motif) ligand 1	1.255277	0.827178

Table S2. Human PCR array data.

Applied Biosystem primer code; Human	Name	FC FINAL	adjPVal
ARTN.Hs00365083_m1	Artemin	2.6461	0.353896
BDNF.Hs00538277_m1	Brain-derived neurotrophic factor	NA	NA
CCL1.Hs00171072_m1	Chemokine (C-C motif) ligand 1	0.9809	0.985747
CCL11.Hs00237013_m1	Chemokine (C-C motif) ligand 11	4.238	0.050675
CCL13.Hs00234646_m1	Chemokine (C-C motif) ligand 12	1.060357	0.869089
CCL14.CCL15.Hs00263137_m1	Chemokine (C-C motif) ligand 14	0.3859	0.279173
CCL16.Hs00171123_m1	Chemokine (C-C motif) ligand 16	NA	NA
CCL17.Hs00171074_m1	Chemokine (C-C motif) ligand 17	0.482628	0.143367
CCL18.Hs00268113_m1	Chemokine (C-C motif) ligand 18	6.602753	6.12E-05
CCL19.Hs00171149_m1	Chemokine (C-C motif) ligand 19	3.581412	2.29E-05
CCL2.Hs00234140_m1	Chemokine (C-C motif) ligand 2	5.134987	1.27E-06
CCL20.Hs00171125_m1	Chemokine (C-C motif) ligand 20	23.54027	0.000138
CCL21.Hs00171076_m1	Chemokine (C-C motif) ligand 21	2.986324	9.96E-06
CCL22.Hs01574247_m1	Chemokine (C-C motif) ligand 22	0.794794	0.374059
CCL23.Hs00270756_m1	Chemokine (C-C motif) ligand 23	1.454472	0.504555
CCL24.Hs00171082_m1	Chemokine (C-C motif) ligand 24	0.955848	0.987641
CCL25.Hs00171144_m1	Chemokine (C-C motif) ligand 25	NA	NA
CCL26.Hs00171146_m1	Chemokine (C-C motif) ligand 26	1.169531	0.595442
CCL27.Hs00171157_m1	Chemokine (C-C motif) ligand 27	0.16084	0.000134
CCL28.Hs00219797_m1	Chemokine (C-C motif) ligand 28	0.474908	0.029053
CCL3.Hs00234142_m1	Chemokine (C-C motif) ligand 3	16.61471	0.000406
CCL4.Hs99999148_m1	Chemokine (C-C motif) ligand 4	2.477342	0.027524
CCL5.Hs00982282_m1	Chemokine (C-C motif) ligand 5	1.342267	0.330106
CCL7.Hs00171147_m1	Chemokine (C-C motif) ligand 7	13.7573	0.000604

CCL8.Hs00271615_m1	Chemokine (C-C motif) ligand 8	18.3125	0.000312
CSF1.Hs00174164_m1	Colony stimulating factor 1 (macrophage)	0.713	0.000466
CSF2.Hs00929873_m1	Colony stimulating factor 2 (granulocyte-macrophage)	0.8536	0.893814
CSF3.Hs99999083_m1	Colony stimulating factor 3 (granulocyte)	25.0016	1.34E-05
CX3CL1.Hs00171086_m1	Chemokine (C-X3-C motif) ligand 1	0.930983	0.859033
CXCL1.Hs00236937_m1	Chemokine (C-X-C motif) ligand 1	18.93007	9.96E-06
CXCL10.Hs00171042_m1	Chemokine (C-X-C motif) ligand 10	6.408928	0.005326
CXCL11.Hs00171138_m1	Chemokine (C-X-C motif) ligand 11	17.9134	0.004117
CXCL12.Hs00930455_m1	Chemokine (C-X-C motif) ligand 12	0.914819	0.708647
CXCL13.Hs00171152_m1	Chemokine (C-X-C motif) ligand 13	NA	NA
CXCL14.Hs00171135_m1	Chemokine (C-X-C motif) ligand 14	0.183163	1.03E-05
CXCL16.Hs00222859_m1	Chemokine (C-X-C motif) ligand 16	1.605064	6.21E-05
CXCL17.Hs01650998_m1	Chemokine (C-X-C motif) ligand 17	8.841316	3.31E-05
CXCL2.Hs00601975_m1	Chemokine (C-X-C motif) ligand 2	11.97302	2.66E-06
CXCL3.Hs00171061_m1	Chemokine (C-X-C motif) ligand 3	13.25804	9.96E-06
CXCL5.Hs00171085_m1	Chemokine (C-X-C motif) ligand 5	82.5431	2.42E-05
CXCL6.Hs00237017_m1	Chemokine (C-X-C motif) ligand 6	5.1695	0.000751
CXCL9.Hs00171065_m1	Chemokine (C-X-C motif) ligand 9	2.997007	0.089332
EBI3.Hs00194957_m1	Epstein-Barr virus induced gene 3	1.634374	0.331126
EDN1.Hs00174961_m1	Endothelin 1	1.544138	0.046452
FGF7.Hs00384281_m1	Fibroblast growth factor 7 or Keratinocyte growth factor	4.295042	3.26E-05
IL10.Hs00961619_m1	Interleukin 10	8.004645	0.000227
IL11.Hs00174148_m1	Interleukin 11	1.602822	0.534722
IL12A.Hs00168405_m1	Interleukin 12 subunit alpha	0.922416	0.26542
IL12B.Hs01011519_m1	Interleukin 12 subunit beta	0.612841	0.670373
IL13.Hs00174379_m1	Interleukin 13	NA	NA
IL15.Hs00174106_m1	Interleukin 15	NA	NA
IL16.Hs00189606_m1	Interleukin 16	0.508381	0.000173
IL17A.Hs00174383_m1	Interleukin 17	NA	NA
IL18.Hs00155517_m1	Interleukin 18	0.226523	0.000277
IL19.Hs00203540_m1	Interleukin 19	NA	NA
IL1A.Hs99999028_m1	Interleukin 1 alpha	5.758932	0.000405
IL1B.Hs99999029_m1	Interleukin 1 beta	10.25314	2.24E-05
IL2.Hs00174114_m1	Interleukin 2	NA	NA
IL20.Hs00218888_m1	Interleukin 20	30.30539	0.00031

IL21.Hs00222327_m1	Interleukin 21	NA	NA
IL22.Hs00220924_m1	Interleukin 22	NA	NA
IL23A.Hs00372324_m1	Interleukin 23 subunit alpha	2.598294	0.048746
IL24.Hs00169533_m1	Interleukin 24	63.731	1.58E-05
IL25.Hs00224471_m1	Interleukin 25	0.2843	0.039623
IL26.Hs00218189_m1	Interleukin 26	NA	NA
IL27.Hs00377366_m1	Interleukin 27	NA	NA
IL28A.Hs00820125_g1	Interleukin 28A	NA	NA
IL29.Hs00601677_g1	Interleukin 29	NA	NA
IL3.Hs00174117_m1	Interleukin 3	NA	NA
IL31.Hs01098710_m1	Interleukin 31	NA	NA
IL32.Hs00170403_m1	Interleukin 32	1.00266	0.988908
IL33.Hs00369211_m1	Interleukin 33	1.177459	0.527756
IL34.Hs00380956_m1	Interleukin 34	0.215375	4.75E-07
IL4.Hs00929862_m1	Interleukin 4	NA	NA
IL5.Hs00174200_m1	Interleukin 5	NA	NA
IL6.Hs00985641_m1	Interleukin 6	54.7143	1.03E-05
IL7.Hs00174202_m1	Interleukin 7	0.44248	0.019158
IL8.Hs99999034_m1	Interleukin 8	71.07711	9.96E-06
IL9.Hs00174125_m1	Interleukin 9	NA	NA
MIF.Hs00236988_g1	Macrophage migration inhibitory factor	1.009424	0.972349
NGF.Hs00171458_m1	Nerve growth factor (beta polypeptide)	1.724755	0.0503
NOS2.Hs00167248_m1	Nitric oxide synthase 2, inducible	0.899343	0.938789
PF4.Hs00236998_m1	Platelet factor 4 (chemokine(C-X-C motif) ligand 4)	2.132868	0.204972
PPBP.Hs00234077_m1	Pro-platelet basic protein (chemokine (C-X-C motif) ligand 7)	3.955725	0.08414
PTGES.Hs00610420_m1	Prostaglandin E synthase	2.452527	0.000123
PTGS2.Hs00153133_m1	Prostaglandin-endoperoxide synthase 2 (cyclooxygenase-2)	5.346755	0.000227
TNF.Hs00174128_m1	Tumor necrosis factor	1.334291	0.112972
TXLNA.Hs00233304_m1	Taxilin alpha	0.830072	0.270347
XCL1.XCL2.Hs00237019_m1	Chemokine (C motif) ligand 1	1.034097	0.972349