

Figure S2

A

| Gene target | IL-1 β ActD 2h | SEM | IL-1 β +ITF2357 ActD 2h | SEM | IL-1 β ActD 5h | SEM | IL-1 β +ITF2357 ActD 5h | SEM |
|-------------|-------------------------|------|-------------------------------------|-----|-------------------------|------|-------------------------------------|-----|
| ADAMTS1 | 48% | 6% | 32% | 4% | 30% | 4% | 18% | 3% |
| ADAMTS13 | 86% | 10% | 76% | 6% | 76% | 6% | 76% | 8% |
| ADORA2A | 98% | 5% | 91% | 6% | 88% | 6% | 79% | 7% |
| ANGPT2 | 135% | 5% | 130% | 24% | 124% | 20% | 99% | 17% |
| BCL2A1 | 103% | 5% | 101% | 10% | 94% | 4% | 88% | 9% |
| BCL2L1 | 84% | 4% | 67% | 4% | 44% | 7% | 29% | 5% |
| BCL2L11 | 29% | 6% | 19% | 2% | 15% | 2% | 12% | 1% |
| BIRC2 | 81% | 2% | 67% | 5% | 53% | 4% | 39% | 1% |
| BIRC5 | 89% | 13% | 84% | 3% | 87% | 13% | 79% | 5% |
| BMP2 | 68% | 2% | 59% | 4% | 34% | 6% | 22% | 3% |
| CCL2 | 78% | 9% | 70% | 12% | 75% | 8% | 59% | 14% |
| CD44 | 104% | 2% | 99% | 3% | 100% | 3% | 99% | 4% |
| CDKN1A | 69% | 5% | 68% | 5% | 53% | 6% | 45% | 5% |
| CDKN1B | 15% | 4% | 8% | 3% | 5% | 1% | 3% | 0% |
| CFLAR | 101% | 4% | 88% | 5% | 82% | 11% | 69% | 6% |
| CSF3 | 67% | 10% | 61% | 21% | 68% | 15% | 55% | 20% |
| CXCL10 | 95% | 10% | 94% | 6% | 87% | 5% | 86% | 6% |
| CXCL11 | 110% | 11% | 116% | 41% | 89% | 12% | 98% | 38% |
| CXCL2 | 28% | 12% | 9% | 4% | 21% | 12% | 5% | 3% |
| CXCL3 | 81% | 6% | 64% | 9% | 81% | 9% | 67% | 16% |
| CXCL5 | 101% | 12% | 105% | 8% | 101% | 18% | 108% | 11% |
| CXCL6 | 107% | 4% | 99% | 2% | 102% | 6% | 108% | 6% |
| CXCL9 | 278% | 207% | 106% | 48% | 302% | 256% | 110% | 54% |
| CXCR4 | 59% | 9% | 69% | 10% | 26% | 8% | 25% | 5% |
| DNMT3B | 66% | 8% | 40% | 3% | 51% | 8% | 33% | 6% |
| EREG | 95% | 7% | 86% | 5% | 81% | 6% | 81% | 9% |
| FASLG | 84% | 10% | 95% | 5% | 92% | 12% | 98% | 5% |
| FGF2 | 89% | 2% | 85% | 6% | 75% | 5% | 71% | 7% |
| FOXO1 | 36% | 9% | 38% | 7% | 9% | 2% | 6% | 1% |
| FOXO3 | 75% | 3% | 72% | 4% | 40% | 3% | 34% | 3% |
| FOXO4 | 82% | 6% | 92% | 4% | 74% | 5% | 76% | 6% |
| GADD45A | 36% | 3% | 25% | 5% | 19% | 2% | 9% | 2% |
| HDAC1 | 96% | 3% | 100% | 3% | 97% | 6% | 102% | 5% |
| HDAC10 | 90% | 8% | 77% | 10% | 78% | 8% | 74% | 10% |
| HDAC11 | 94% | 10% | 98% | 5% | 85% | 9% | 92% | 5% |
| HDAC2 | 82% | 12% | 68% | 6% | 71% | 13% | 65% | 7% |
| HDAC3 | 94% | 5% | 89% | 5% | 91% | 6% | 86% | 3% |
| HDAC4 | 81% | 6% | 83% | 2% | 63% | 6% | 61% | 3% |
| HDAC5 | 81% | 7% | 66% | 3% | 73% | 7% | 58% | 6% |
| HDAC6 | 88% | 3% | 88% | 6% | 79% | 5% | 73% | 7% |
| HDAC7 | 73% | 8% | 63% | 5% | 51% | 7% | 41% | 7% |
| HDAC8 | 92% | 2% | 90% | 3% | 85% | 5% | 73% | 3% |
| HDAC9 | 101% | 7% | 89% | 4% | 70% | 10% | 74% | 7% |
| ICAM1 | 90% | 2% | 91% | 5% | 82% | 5% | 76% | 5% |
| IFNB1 | 97% | 15% | 84% | 18% | 66% | 11% | 56% | 19% |
| IFNGR1 | 91% | 12% | 81% | 11% | 69% | 5% | 64% | 7% |
| IFNGR2 | 111% | 10% | 93% | 12% | 99% | 11% | 99% | 9% |
| IL1A | 78% | 13% | 64% | 7% | 56% | 15% | 44% | 13% |
| IL1B | 81% | 12% | 73% | 15% | 80% | 19% | 77% | 26% |
| IL1F5 | 91% | 6% | 73% | 12% | 74% | 1% | 77% | 8% |
| IL1F9 | 69% | 11% | 132% | 76% | 43% | 15% | 77% | 19% |
| IL1RN | 88% | 5% | 122% | 26% | 95% | 14% | 72% | 18% |
| IL6 | 56% | 15% | 31% | 14% | 51% | 17% | 26% | 14% |
| IL8 | 70% | 9% | 53% | 15% | 71% | 8% | 49% | 16% |
| IRAK2 | 57% | 10% | 46% | 10% | 27% | 9% | 16% | 5% |
| IRF1 | 17% | 4% | 15% | 4% | 4% | 1% | 3% | 0% |
| ITGA2 | 73% | 6% | 69% | 13% | 61% | 6% | 83% | 12% |
| LAMB3 | 94% | 2% | 92% | 6% | 83% | 7% | 89% | 4% |
| MMP1 | 96% | 5% | 103% | 9% | 88% | 7% | 105% | 9% |
| MMP10 | 116% | 15% | 107% | 7% | 108% | 17% | 102% | 5% |
| MMP13 | 141% | 49% | 107% | 45% | 99% | 40% | 105% | 28% |
| MMP3 | 100% | 3% | 95% | 2% | 97% | 2% | 104% | 3% |
| MMP7 | 84% | 10% | 114% | 7% | 83% | 4% | 103% | 6% |
| MMP8 | 254% | 200% | 106% | 40% | 288% | 217% | 115% | 74% |
| PTGES | 102% | 3% | 101% | 2% | 107% | 6% | 102% | 5% |

| | | | | | | | | |
|----------|------|-----|------|-----|------|-----|------|-----|
| MYD88 | 94% | 5% | 88% | 4% | 78% | 7% | 70% | 4% |
| NCAM1 | 87% | 7% | 130% | 35% | 73% | 9% | 116% | 41% |
| NFKB1 | 98% | 2% | 93% | 4% | 72% | 4% | 58% | 7% |
| NFKBIA | 2% | 0% | 1% | 0% | 1% | 0% | 0% | 0% |
| NOS2 | 85% | 8% | 82% | 13% | 72% | 6% | 57% | 17% |
| PDGFB | 74% | 10% | 47% | 12% | 23% | 8% | 12% | 4% |
| PTGS2 | 75% | 13% | 41% | 13% | 68% | 16% | 44% | 19% |
| RHOB | 23% | 3% | 20% | 2% | 3% | 1% | 1% | 0% |
| SELE | 65% | 11% | 44% | 10% | 46% | 9% | 36% | 8% |
| SERPINA1 | 100% | 5% | 102% | 4% | 102% | 10% | 105% | 7% |
| SOCS3 | 7% | 1% | 4% | 0% | 2% | 0% | 2% | 0% |
| SOD2 | 100% | 4% | 97% | 4% | 98% | 2% | 93% | 2% |
| TIMP3 | 131% | 83% | 21% | 10% | 73% | 20% | 36% | 18% |
| TLR1 | 73% | 5% | 80% | 6% | 53% | 6% | 61% | 9% |
| TLR2 | 111% | 10% | 102% | 15% | 98% | 11% | 89% | 13% |
| TLR4 | 86% | 5% | 76% | 3% | 71% | 5% | 52% | 3% |
| TNF | 23% | 13% | 8% | 3% | 12% | 9% | 3% | 1% |
| VCAM1 | 101% | 3% | 99% | 5% | 95% | 6% | 94% | 8% |
| B2M | 106% | 5% | 100% | 1% | 109% | 3% | 104% | 3% |
| HPRT1 | 102% | 2% | 106% | 4% | 97% | 4% | 106% | 2% |
| RPL13A | 96% | 1% | 94% | 2% | 92% | 2% | 92% | 2% |
| GAPDH | 97% | 3% | 100% | 7% | 103% | 6% | 99% | 3% |
| ACTB | 100% | 3% | 101% | 3% | 101% | 2% | 100% | 4% |

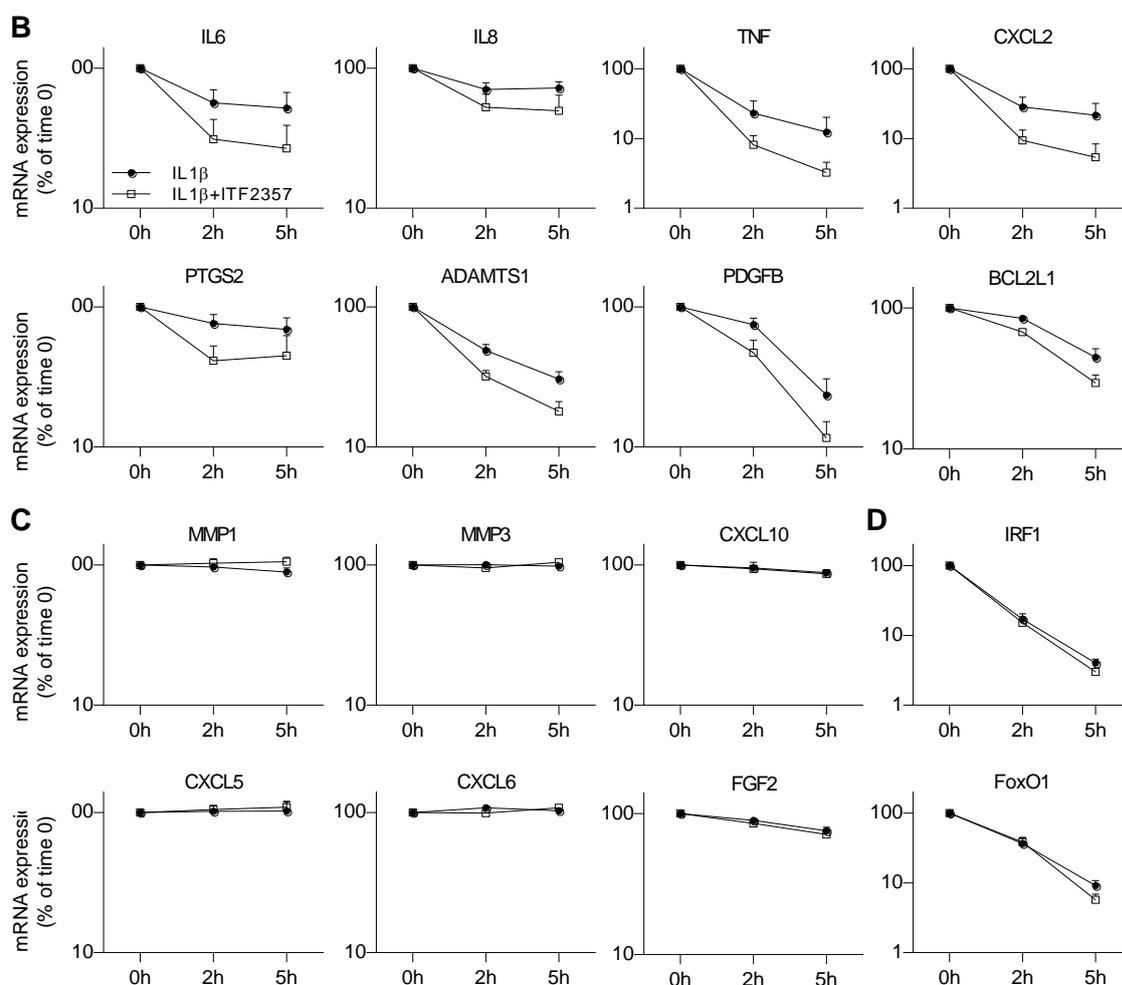


Figure S2. Effects of ITF2357 on mRNA stability. (A-B) RA FLS (n=4) were left untreated or were treated with ITF2357 prior to incubation with IL-1 β for 2h. Transcription was then blocked with 10 μ g/ml of actinomycin D (ActD) and RNA extracted at the indicated time points from the start of ActD treatment. mRNA degradation in the presence or absence of ITF2357 was monitored using customized qPCR arrays. mRNA expression values for 0h time point were normalized to 100%. (A) Mean percentage \pm SEM of remaining mRNA after 2h and 5h ActD treatment, calculated relative to 100% value of relative controls from 0h ActD treatment. (B-C-D) RA FLS (n=4) were treated as in A. Representative genes displaying enhanced mRNA degradation by ITF2357 (B), no appreciable decay regardless of IL-1 β stimulation (C), or no destabilization by ITF2357 (D).