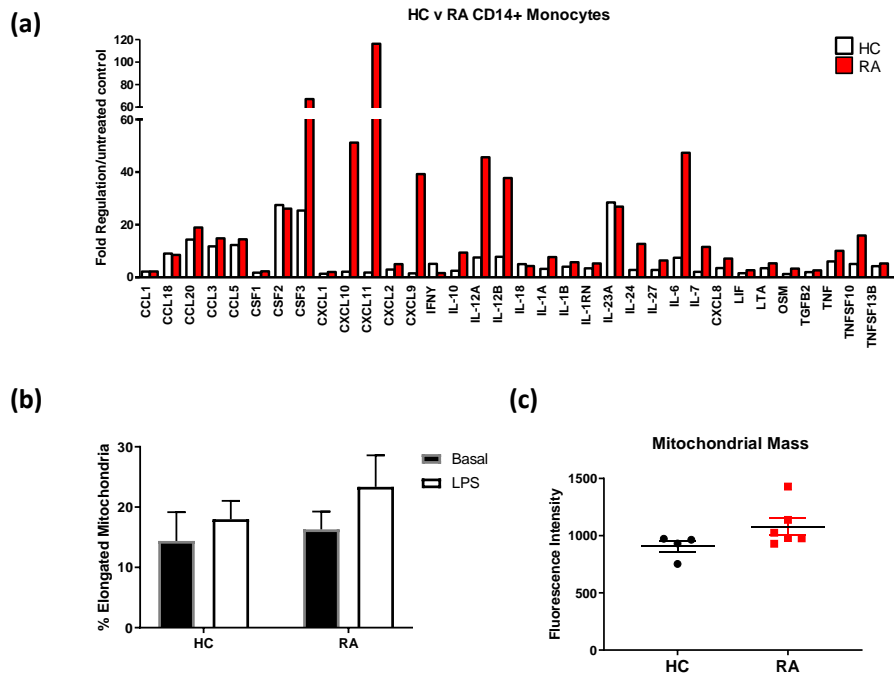


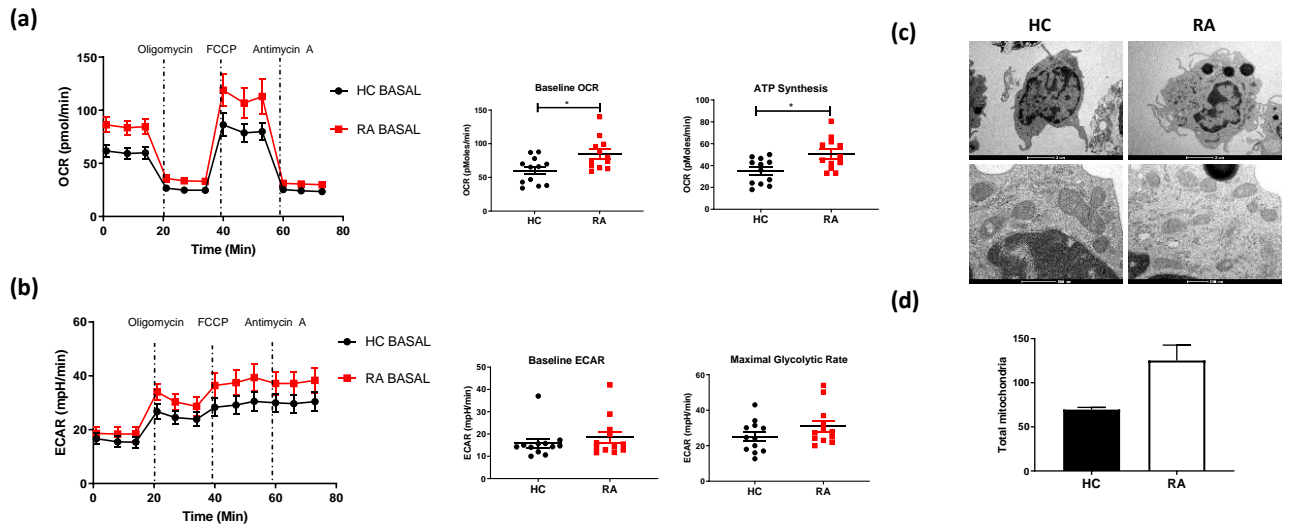
Gene Name	Forward Primer	Reverse Primer
<i>RPLPO</i>	5' GCGTCCTCGTGGAAGTGACATCG 3'	5' TCAGGGATTGCCACGCAGGG 3'
<i>HPRT1</i>	5' ATGGACAGGACTGAACGTCTTG 3'	5' GGCTACAATGTGATGGCCTC 3'
<i>TNFα</i>	5' ACCTCTCTCTAATCAGCCCTC 3'	5' GGTTGAGAAGATGATCTGACTG 3'
<i>IL-6</i>	5' CCCTGAGAAAGGAGACATGTAAC 3'	5' CCTCTTTGCTGCTTTCACACATG 3'
<i>IL-1β</i>	5' CTCAAGTGTCTGAAGCAGCCAT 3'	5' CATCATTTCACTGGCGAGCTCA 3'
<i>OSM</i>	5' ACTCCTGGACCCCTATATACG 3'	5' AGTGCTCTCTCAGTTTAGGAACAT 3'
<i>CXCL10</i>	5' TTCAAGGAGTACCTCTCTAGAA 3'	5' GGTTGATTACTAATGCTGATGCAG 3'
<i>CXCL11</i>	5' GGCTTCCCATGTTCAAAGAG 3'	5' TCTCAATATCTGCCACTTTCCTG 3'
<i>IL-27</i>	5' CTTTGC GGAATCTCACCTGCC 3'	5' AGGGAAACATCAGGGAGCTGC 3'
<i>IL-8</i>	5' TTGGCAGCCTTCTGATTC 3'	5' TGGCAAACTGCACCTTCAC 3'
<i>PFKFB3</i>	5' ACCAAAGATCACCCACGGATGT 3'	5' AGCGAGTGCAGAATGGACACAA 3'
<i>HK2</i>	5' TTCTGTCTCAGATTGAGAGTGAC 3'	5' TTGCAGGATGGCTCGGACTTG 3'
<i>HIF1α</i>	5' GAAACTTCTGGATGCTGGTGATT 3'	5' GCAATTCATCTGTGCTTTCATGTCA 3'
<i>STAT3</i>	5' TTCACTTGGGTGGAGAAG 3'	5' CGGACTGGATCTGGGTCT 3'
<i>IL-1α</i>	5' CAAAGAAGTCAAGATGGCCAA 3'	5' CTGTAACAGTTCTTCAGGTCT 3'

Supplementary table 1: List of designed primers for real-time PCR analysis



Supplementary figure 1:

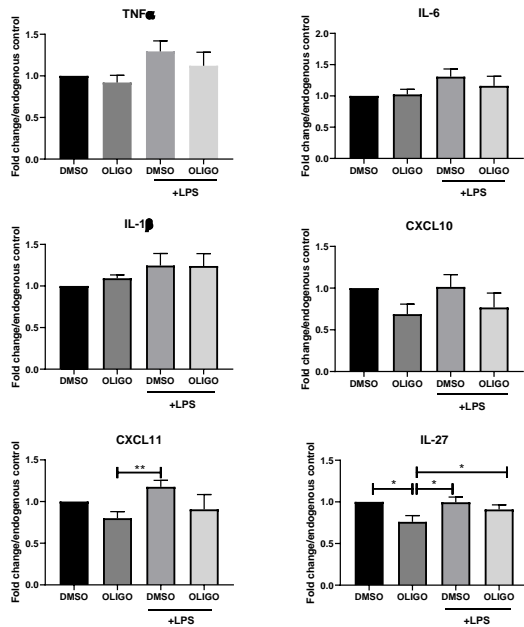
(a) Bar graph demonstrating 34 cytokine/chemokine genes upregulated in response to 3 hr LPS stimulation (100 ng mL^{-1}) in healthy and RA CD14⁺ monocytes (n=3). **(b)** Gating strategy for identification of CD14⁺ monocytes in blood. **(c)** Bar graphs demonstrating the frequency of elongated mitochondria as a percentage of total mitochondria present in HC (n=3) and RA (n=3) CD14⁺ monocytes +/- LPS. Data expressed as mean \pm SEM.



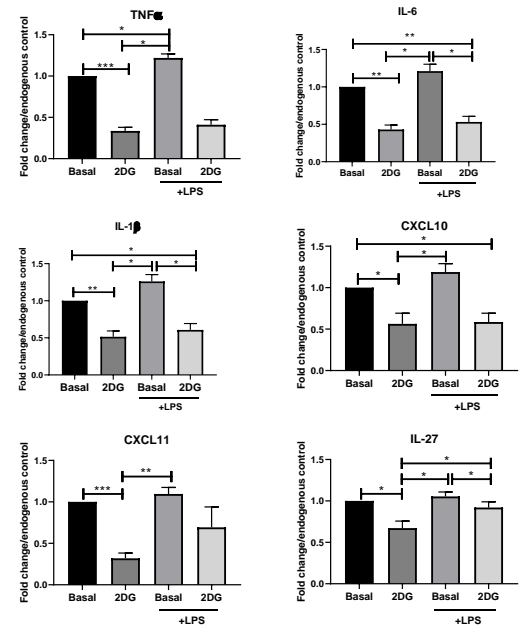
Supplementary figure 2:

(a) Seahorse bioenergetics profile (average) OCR of healthy (n=12) and RA (n=12) monocytes before and after injections of oligomycin, FCCP and antimycin A in *ex vivo* resting monocytes with dot plots depicting baseline OCR and ATP synthesis. **(b)** Seahorse bioenergetics profile (average) ECAR of healthy (n=12) and RA (n=12) monocytes before and after injections of oligomycin, FCCP and antimycin A in *ex vivo* resting monocytes with dot plots depicting baseline ECAR and Maximal Glycolytic Rate. **(c)** Representative TEM images of healthy control and RA *ex vivo* resting monocytes. Scale bar represents 2 μm and 500 nm. **(d)** Bar graphs represent quantification of total mitochondria present in HC (n=3) and RA (n=3) CD14⁺ resting monocytes. Data expressed as mean \pm SEM, Mann-Whitney *U*-test and two-way ANOVA used as appropriate, **P* < 0.05.

(a)

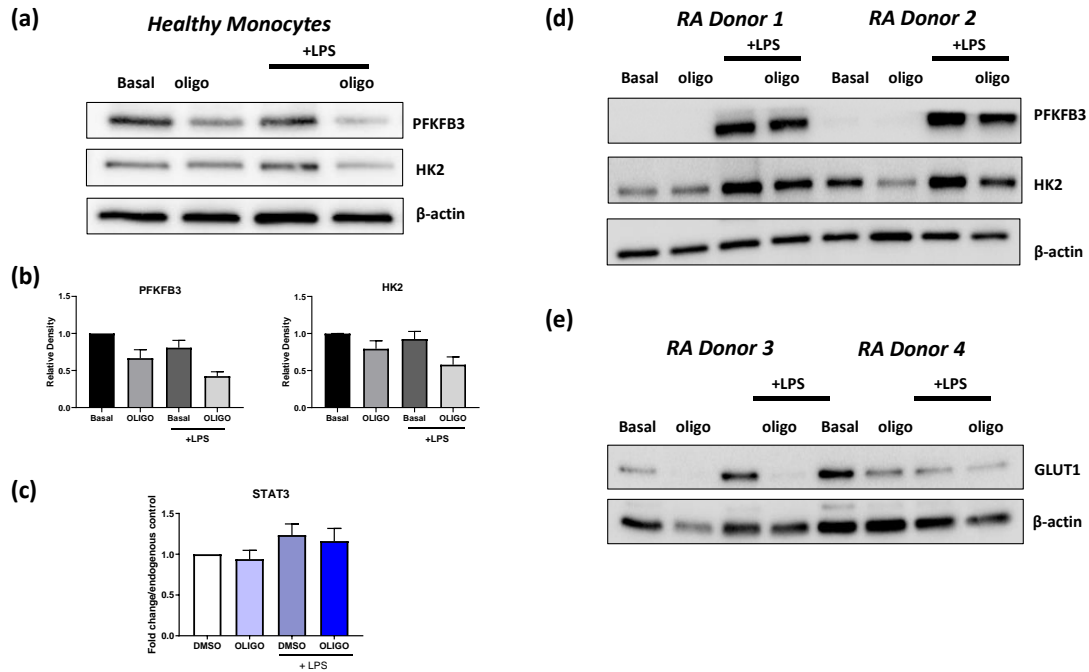


(b)



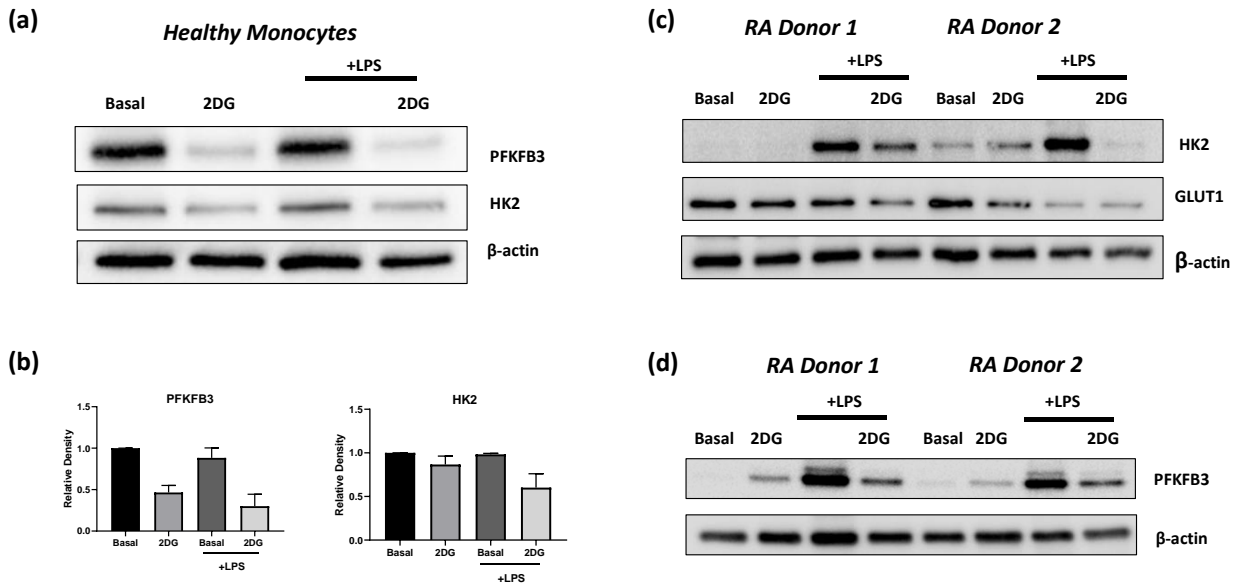
Supplementary figure 3:

HC monocytes stimulated with LPS (100 ng mL⁻¹) for 3 hr +/- (a) oligomycin (n=5) and (b) 2DG (n=4 or 5). Gene expression of *TNF α* , *IL-6*, *IL-1 β* , *CXCL10*, *CXCL11* and *IL-27*. Data expressed as mean \pm SEM, paired t-test used, * P < 0.05, ** P < 0.01, *** P < 0.005.



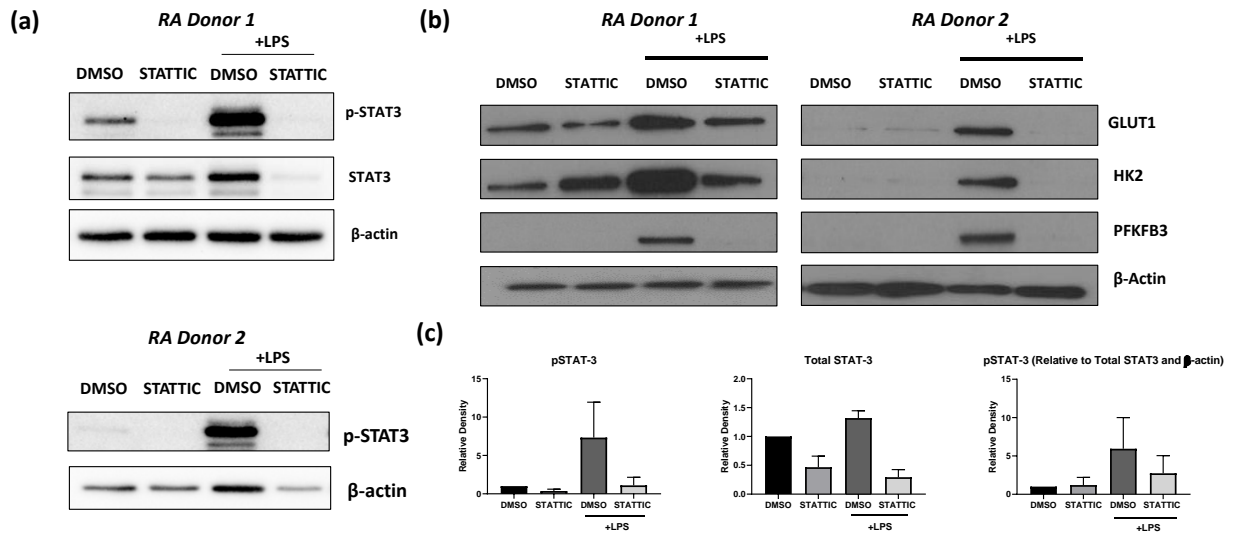
Supplementary figure 4:

Representative western blots demonstrating HK2 and PFKFB3 protein expression in HC CD14⁺ monocytes stimulated with LPS (100 ng mL⁻¹) for 3 hr +/- **(a)** oligomycin and **(c)** 2DG. Bar graphs representing densitometry quantification of PFKFB3 and HK2 normalized to β -actin in HC CD14⁺ monocytes stimulated with LPS (100 ng/mL) for 3 hr +/- **(b)** oligomycin (n=4) and **(d)** 2DG (n=4). Data expressed as mean \pm SEM.



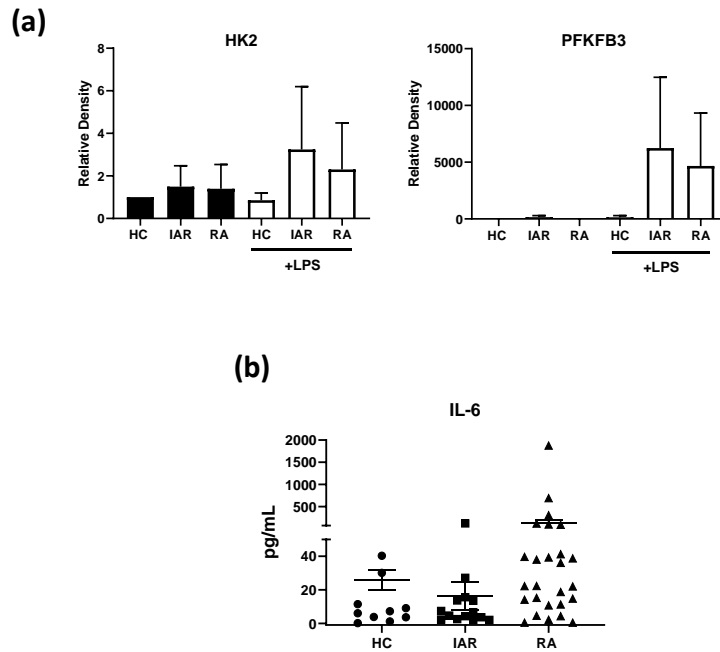
Supplementary figure 5:

Western blots demonstrating HK2 and PFKFB3 protein expression in two independent RA CD14⁺ patient monocytes stimulated with LPS (100 ng mL⁻¹) for 3 hr +/- **(a)** oligomycin (n=2) and **(b)** 2DG (n=2). **(c)** *STAT3* gene expression in response to LPS +/- oligomycin in RA CD14⁺ monocytes (n=7). Data represented as mean \pm SEM.



Supplementary figure 6:

(a) Western blots demonstrating pSTAT3 protein expression in RA CD14⁺ monocytes (n=2) stimulated with LPS (100 ng mL⁻¹) for 3 hr +/- STATTIC. **(b)** Bar graphs representing densitometry quantification of pSTAT3 normalized to β -actin in RA CD14⁺ monocytes (n=2) stimulated with LPS (100 ng mL⁻¹) for 3 hr +/- STATTIC. **(c)** Western blots demonstrating HK2, PFKFB3 and GLUT1 protein expression in two independent RA CD14⁺ patient monocytes stimulated with LPS (100 ng mL⁻¹) for 3 hr +/- STATTIC (n=2). Data expressed as mean \pm SEM.



Supplementary Figure 7:

(a) Bar graphs representing densitometry quantification of HK2 and PFKFB3 normalized to β -actin in HC (n=2), IAR (n=2) and RA (n=2) CD14⁺ monocytes following activation with LPS. **(b)** Dot plot representing expression of secreted IL-6 in HC (n=15), IAR (n=14) and RA (n=31) patient serums.