

Additional file 5: Table S4 List of significantly upregulated and downregulated gene sets in PBLs of sarcoidosis patients at FDR < 1% threshold. Note that several T cell-associated pathways were downregulated in PBLs of sarcoidosis patients.

| Upregulated gene set | Number of genes | FDR |
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| REACTOME_INTERFERON_GAMMA_SIGNALING | 56 | 0 |
| REACTOME_INTERFERON_ALPHA_BETA_SIGNALING | 62 | 0 |
| REACTOME_INTERFERON_SIGNALING | 149 | 0 |
| REACTOME_CYTOKINE_SIGNALING_IN_IMMUNE_SYSTEM | 258 | 0 |
| REACTOME_INNATE_IMMUNE_SYSTEM | 246 | 0 |
| KEGG_LEISHMANIA_INFECTION | 69 | 0 |
| KEGG_NEUROACTIVE_LIGAND_RECEPTOR_INTERACTION | 267 | 0 |
| BIOCARTA_COMP_PATHWAY | 18 | 0 |
| REACTOME_COMPLEMENT_CASCADE | 29 | 0 |
| KEGG_COMPLEMENT_AND_COAGULATION_CASCADES | 68 | 0 |
| REACTOME_ACTIVATED_TAK1_MEDIATES_P38_MAPK_ACTIVATION | 16 | 0 |
| REACTOME_EICOSANOID_LIGAND_BINDING_RECEPTORS | 14 | 0.0003 |
| REACTOME_TOLL_RECEPTOR_CASCADES | 112 | 0.0006 |
| REACTOME_ACTIVATED_TLR4_SIGNALING | 89 | 0.0006 |
| REACTOME_ANTIGEN_PROCESSING_CROSS_PRESENTATION | 71 | 0.0014 |
| PID_UPA_UPAR_PATHWAY | 40 | 0.0017 |
| KEGG_TOLL_LIKE_RECEPTOR_SIGNALING_PATHWAY | 100 | 0.0017 |
| REACTOME_TAK1_ACTIVATES_NFKB_BY_PHOSPHORYLATION_AND_ACTIVATION_OF_IKKS_COMPLEX | 23 | 0.0017 |
| KEGG_NOD_LIKE_RECEPTOR_SIGNALING_PATHWAY | 61 | 0.0017 |
| REACTOME_TRIF_MEDIATED_TLR3_SIGNALING | 71 | 0.0018 |
| KEGG_SYSTEMIC_LUPUS_ERYTHEMATOSUS | 90 | 0.0018 |
| PID_IFNG_PATHWAY | 39 | 0.0019 |
| REACTOME_MYD88_MAL_CASCADE_INITIATED_ON_PLASMA_MEMBRANE | 79 | 0.0019 |
| BIOCARTA_CLASSIC_PATHWAY | 13 | 0.0019 |
| REACTOME_IL1_SIGNALING | 38 | 0.0019 |
| REACTOME_TRAF6_MEDIATED_INDUCTION_OF_NFKB_AND_MAP_KINASES_UPON_TLR7_8_OR_9_ACTIVATION | 73 | 0.0025 |
| KEGG_NOTCH_SIGNALING_PATHWAY | 47 | 0.0029 |
| REACTOME_REGULATION_OF_IFNA_SIGNALING | 24 | 0.0030 |
| PID_INTEGRIN_A9B1_PATHWAY | 25 | 0.0030 |
| REACTOME_ACTIVATED_NOTCH1_TRANSMITS_SIGNAL_TO_THE_NUCLEUS | 26 | 0.0031 |
| REACTOME_GPCR_LIGAND_BINDING | 378 | 0.0031 |
| REACTOME_NFKB_AND_MAP_KINASES_ACTIVATION_MEDIATED_BY_TLR4_SIGNALING_REPERTOIRE | 69 | 0.0035 |
| REACTOME_RIG_I_MDA5_MEDIATED_INDUCTION_OF_IFN_ALPHA_BETA_PATHWAYS | 71 | 0.0041 |
| REACTOME_TRAF6_MEDIATED_IRF7_ACTIVATION | 30 | 0.0046 |
| REACTOME_DEGRADATION_OF_THE_EXTRACELLULAR_MATRIX | 29 | 0.0047 |
| KEGG_DORSO_VENTRAL_AXIS_FORMATION | 23 | 0.0053 |
| REACTOME_SIGNALING_BY_ILS | 105 | 0.0053 |
| REACTOME_NUCLEOTIDE_BINDING_DOMAIN_LEUCINE_RICH_REPEAT_CONTAINING_RECEPTOR_NLR_SIGNALING_PATHWAYS | 43 | 0.0056 |
| REACTOME_CHONDROITIN_SULFATE_DERMATAN_SULFATE_METABOLISM | 47 | 0.0056 |
| REACTOME_GABA_B_RECEPTOR_ACTIVATION | 38 | 0.0066 |
| REACTOME_CHONDROITIN_SULFATE_BIOSYNTHESIS | 19 | 0.0067 |
| REACTOME_GROWTH_HORMONE_RECEPTOR_SIGNALING | 24 | 0.0067 |
| REACTOME_RIP_MEDIATED_NFKB_ACTIVATION_VIA_DAI | 18 | 0.0067 |
| REACTOME_INITIAL_TRIGGERING_OF_COMPLEMENT | 13 | 0.0069 |
| REACTOME_G_ALPHA_I_SIGNALING_EVENTS | 184 | 0.0082 |
| REACTOME_AMYLOIDS | 47 | 0.0085 |
| REACTOME_EXTRACELLULAR_MATRIX_ORGANIZATION | 85 | 0.0088 |
| Downregulated gene set | Number of genes | FDR |
| REACTOME_TRANSLATION | 142 | 0 |
| REACTOME_3_UTR_MEDIATED_TRANSLATIONAL_REGULATION | 102 | 0 |
| REACTOME_SRP_DEPENDENT_COTRANSLATIONAL_PROTEIN_TARGETING_TO_MEMBRANE | 105 | 0 |
| REACTOME_PEPTIDE_CHAIN_ELONGATION | 82 | 0 |
| KEGG_RIBOSOME | 84 | 0 |
| REACTOME_NONSENSE_MEDIATED_DECAY_ENHANCED_BY_THE_EXON_JUNCTION_COMPLEX | 102 | 0 |
| REACTOME_INFLUENZA_VIRAL_RNA_TRANSCRIPTION_AND_REPLICATION | 98 | 0 |
| REACTOME_INFLUENZA_LIFE_CYCLE | 131 | 0 |
| REACTOME_METABOLISM_OF_RNA | 247 | 0 |
| REACTOME_ACTIVATION_OF_THE_MRNA_UPON_BINDING_OF_THE_CAP_BINDING_COMPLEX_AND_EIFS_AND_SUBSEQUENT_BINDING_TO_43S | 54 | 0 |
| REACTOME_FORMATION_OF_THE_TERNARY_COMPLEX_AND_SUBSEQUENTLY_THE_43S_COMPLEX | 46 | 0 |
| REACTOME_METABOLISM_OF_MRNA | 203 | 0 |
| REACTOME_PROCESSING_OF_CAPPED_INTRON_CONTAINING_PRE_MRNA | 133 | 0 |
| REACTOME_MRNA_PROCESSING | 153 | 0 |
| BIOCARTA_TCR_PATHWAY | 42 | 0 |
| REACTOME_METABOLISM_OF_NON_CODING_RNA | 47 | 0 |

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| REACTOME_METABOLISM_OF_PROTEINS | 408 | 0 |
| REACTOME_GENERIC_TRANSCRIPTION_PATHWAY | 327 | 0.0000 |
| REACTOME_MITOCHONDRIAL_TRNA_AMINOACYLATION | 21 | 0.0000 |
| REACTOME_MRNA_SPLICING | 104 | 0.0000 |
| REACTOME_MITOCHONDRIAL_PROTEIN_IMPORT | 47 | 0.0001 |
| KEGG_RNA_POLYMERASE | 26 | 0.0002 |
| KEGG_PROTEIN_EXPORT | 23 | 0.0002 |
| REACTOME_TRNA_AMINOACYLATION | 42 | 0.0003 |
| KEGG_AMINOACYL_TRNA_BIOSYNTHESIS | 41 | 0.0003 |
| REACTOME_CLEAVAGE_OF_GROWING_TRANSCRIPT_IN_THE_TERMINATION_REGION_ | 40 | 0.0004 |
| REACTOME_TRANSPORT_OF_MATURE_TRANSCRIPT_TO_CYTOPLASM | 50 | 0.0004 |
| KEGG_SPLICEOSOME | 118 | 0.0004 |
| REACTOME_INTERACTIONS_OF_VPR_WITH_HOST_CELLULAR_PROTEINS | 32 | 0.0004 |
| PID_CD8_TCR_PATHWAY | 53 | 0.0006 |
| KEGG_T_CELL_RECEPTOR_SIGNALING_PATHWAY | 107 | 0.0007 |
| REACTOME_RESPIRATORY_ELECTRON_TRANSPORT_ATP_SYNTHESIS_BY_CHEMIOSMOTIC_COUPLING_AND_HEAT_PRODUCTION_BY_UNCOUPLING_PROTEINS_ | 85 | 0.0009 |
| BIOCARTA_TCRA_PATHWAY | 11 | 0.0009 |
| REACTOME_RNA_POL_II_TRANSCRIPTION | 95 | 0.0009 |
| REACTOME_PURINE_RIBONUCLEOSIDE_MONOPHOSPHATE_BIOSYNTHESIS | 11 | 0.0010 |
| KEGG_RNA_DEGRADATION | 57 | 0.0015 |
| REACTOME_BIOSYNTHESIS_OF_THE_N_GLYCAN_PRECURSOR_DOLICHOL_LIPID_LINKED_OLIGOSACCHARIDE_LLO_AND_TRANSFER_TO_A_NASCENT_PROTEIN | 26 | 0.0016 |
| REACTOME_SLBP_DEPENDENT_PROCESSING_OF_REPLICATION_DEPENDENT_HISTONE_PRE_MRNAS | 11 | 0.0024 |
| BIOCARTA_STATHMIN_PATHWAY | 19 | 0.0033 |
| REACTOME_HIV_LIFE_CYCLE | 110 | 0.0044 |
| REACTOME_NEP_NS2_INTERACTS_WITH_THE_CELLULAR_EXPORT_MACHINERY | 27 | 0.0046 |
| BIOCARTA_GPCR_PATHWAY | 32 | 0.0046 |
| REACTOME_MRNA_3_END_PROCESSING | 31 | 0.0047 |
| BIOCARTA_CTLA4_PATHWAY | 19 | 0.0050 |
| REACTOME_RESPIRATORY_ELECTRON_TRANSPORT | 68 | 0.0051 |
| REACTOME_TCA_CYCLE_AND_RESPIRATORY_ELECTRON_TRANSPORT | 121 | 0.0052 |
| REACTOME_GENERATION_OF_SECOND_MESSENGER_MOLECULES | 24 | 0.0059 |
| BIOCARTA_TOB1_PATHWAY | 19 | 0.0059 |
| REACTOME_TRANSPORT_OF_MATURE_MRNA_DERIVED_FROM_AN_INTRONLESS_TRANSCRIPT | 32 | 0.0065 |
| REACTOME_PROCESSING_OF_CAPPED_INTRONLESS_PRE_MRNA | 23 | 0.0069 |
| REACTOME_TCR_SIGNALING | 49 | 0.0072 |
| REACTOME_LATE_PHASE_OF_HIV_LIFE_CYCLE | 97 | 0.0079 |
| BIOCARTA_CSK_PATHWAY | 22 | 0.0087 |
| REACTOME_REGULATORY_RNA_PATHWAYS | 21 | 0.0088 |