

Table S5

Gene networks implicated in Multiple Sclerosis (MS) pathogenesis from miR-20a knock-in and -down experiments and from mRNA expression in whole blood

| Pathway | P Value | No. Genes Represented | No. Genes on Pathway | No. Genes in miRNA list |
|--|----------|-----------------------|----------------------|-------------------------|
| miR-20a Jurkat differential expression, miR-20a target genes, and MS mRNA differential expression | | | | 31 |
| Transcription_Sin3 and NuRD in transcription regulation | 3.11E-06 | 4 | 150 | |
| Immune response_Antiviral actions of Interferons | 3.46E-06 | 4 | 154 | |
| Immune response_TLR3 and TLR4 induce TICAM1-specific signaling pathway | 2.85E-05 | 3 | 88 | |
| Immune response_IFN alpha/beta signaling pathway | 3.36E-05 | 3 | 93 | |
| Immune response_Innate immunity response to RNA viral infection | 7.92E-05 | 3 | 124 | |
| Signal transduction_Activin A signaling regulation | 4.06E-04 | 3 | 216 | |
| G-protein signaling_Ras family GTPases in kinase cascades (scheme) | 1.66E-03 | 2 | 90 | |
| Cell cycle_Chromosome condensation in prometaphase | 1.84E-03 | 2 | 95 | |
| Apoptosis and survival_Cytoplasmic/mitochondrial transport of proapoptotic proteins Bid, Bmf and Bim | 1.92E-03 | 2 | 97 | |
| Cell cycle_Sister chromatid cohesion | 2.08E-03 | 2 | 101 | |
| Regulation of lipid metabolism_Regulation of lipid metabolism via LXR, NF-Y and SREBP | 2.25E-03 | 2 | 105 | |
| Development_Glucocorticoid receptor signaling | 2.78E-03 | 2 | 117 | |
| Transcription_Ligand-dependent activation of the ESR1/SP pathway | 3.62E-03 | 2 | 134 | |
| Transcription_Role of heterochromatin protein 1 (HP1) family in transcriptional silencing | 3.67E-03 | 2 | 135 | |
| Cell cycle_Initiation of mitosis | 4.00E-03 | 2 | 141 | |
| Cell cycle_Role of Nek in cell cycle regulation | 5.24E-03 | 2 | 162 | |
| miR-20a Jurkat differential expression and MS mRNA differential expression | | | | 93 |
| Translation _Regulation of translation initiation | 6.16E-14 | 14 | 376 | |
| Signal transduction_Activin A signaling regulation | 1.15E-12 | 11 | 216 | |
| Translation_(L)-selenoaminoacids incorporation in proteins during translation | 1.05E-11 | 12 | 349 | |
| Vitamin B7 (biotin) metabolism | 1.69E-09 | 7 | 99 | |
| Immune response_Antiviral actions of Interferons | 8.89E-07 | 6 | 154 | |
| Transcription_Sin3 and NuRD in transcription regulation | 1.60E-05 | 5 | 150 | |
| Immune response_TLR3 and TLR4 induce TICAM1-specific signaling pathway | 8.28E-04 | 3 | 88 | |
| Immune response_IFN alpha/beta signaling pathway | 9.72E-04 | 3 | 93 | |
| Immune response_Innate immunity response to RNA viral infection | 2.22E-03 | 3 | 124 | |
| Transcription_Ligand-dependent activation of the ESR1/SP pathway | 2.76E-03 | 3 | 134 | |
| Immune response _Immunological synapse formation | 3.32E-03 | 4 | 294 | |

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|--|----------|----|-----|-------------|
| Development_NOTCH1-mediated pathway for NF-KB activity modulation | 5.03E-03 | 3 | 166 | |
| Development_Notch Signaling Pathway | 6.10E-03 | 3 | 178 | |
| Transport_Macropinocytosis regulation by growth factors | 8.50E-03 | 3 | 201 | |
| miR-20a Jurkat differential expression and miR-20a target genes | | | | 914 |
| Cholesterol Biosynthesis | 7.23E-05 | 7 | 63 | |
| Regulation of lipid metabolism_RXR-dependent regulation of lipid metabolism via PPAR, RAR and VDR | 7.10E-04 | 6 | 66 | |
| DNA damage_NHEJ mechanisms of DSBs repair | 1.05E-03 | 6 | 71 | |
| Transcription_Ligand-dependent activation of the ESR1/SP pathway | 1.61E-03 | 8 | 134 | |
| Regulation of lipid metabolism_Regulation of lipid metabolism via LXR, NF-Y and SREBP | 1.66E-03 | 7 | 105 | |
| Cell cycle_Start of DNA replication in early S phase | 4.04E-03 | 7 | 123 | |
| Apoptosis and survival_Cytoplasmic/mitochondrial transport of proapoptotic proteins Bid, Bmf and Bim | 5.07E-03 | 6 | 97 | |
| miR-20a Jurkat differential expression | | | | 2741 |
| Cholesterol Biosynthesis | 1.14E-05 | 11 | 63 | |
| Translation _Regulation of translation initiation | 1.90E-04 | 28 | 376 | |
| Regulation of lipid metabolism_Regulation of lipid metabolism via LXR, NF-Y and SREBP | 1.21E-03 | 11 | 105 | |
| Translation_Insulin regulation of translation | 2.21E-03 | 14 | 165 | |
| Cell cycle_Spindle assembly and chromosome separation | 2.56E-03 | 19 | 262 | |
| Translation_(L)-selenoaminoacids incorporation in proteins during translation | 3.30E-03 | 23 | 349 | |
| Development_Fit3 signalling | 3.84E-03 | 13 | 157 | |