## Default miRComb output

/home/mvila/Baixades/TCGA/stomach
May 13, 2015

## 1 Exploratory analysis of miRNA dataset

$$
\begin{array}{ll}
\hline \text { Number of miRNAs analysed } & 330 \\
\hline \text { Number of samples } & 443
\end{array}
$$

Table 1: Basic information of the miRNA dataset.

|  | group.n | CvH | center | sample | batch |
| :--- | :--- | :--- | :--- | :--- | :--- |
| 1 | NT: 37 | Min. $: 0.0000$ | BR :138 | TCGA-3M-AB46-01: 1 | Batch 220:58 |
| 2 | TP:406 | 1st Qu.:1.0000 | VQ $: 66$ | TCGA-3M-AB47-01: 1 | Batch 427:58 |
| 3 |  | Median $: 1.0000$ | CG $: 43$ | TCGA-B7-5816-01: 1 | Batch 269:52 |
| 4 |  | Mean :0.9165 | HU $: 41$ | TCGA-B7-5818-01:1 | Batch 57:31 |
| 5 |  | 3rd Qu.:1.0000 | D7:40 | TCGA-B7-A5TI-01:1 | Batch 95:29 |
| 6 |  | Max. :1.0000 | CD :27 | TCGA-B7-A5TJ-01:1 | Batch 242:28 |
| 7 |  |  | (Other):88 | (Other) $: 437$ | (Other) $: 187$ |

Table 2: Summary of the phenotypical information of the miRNA dataset.


Figure 1: PCA and density plot for miRNAs.

## 2 Exploratory analysis of mRNA dataset

| Number of mRNAs analysed | 18565 |
| :--- | :--- |
| Number of samples | 443 |

Table 3: Basic information of the mRNA dataset.

|  | group.n | CvH | center | sample | batch |
| :--- | :--- | :--- | :--- | :--- | :--- |
| 1 | NT: 37 | Min. :0.0000 | BR :138 | TCGA-3M-AB46-01: 1 | Batch 220:58 |
| 2 | TP:406 | 1st Qu.:1.0000 | VQ $: 66$ | TCGA-3M-AB47-01:1 | Batch 427:58 |
| 3 |  | Median $: 1.0000$ | CG $: 43$ | TCGA-B7-5816-01:1 | Batch 269:52 |
| 4 |  | Mean :0.9165 | HU $: 41$ | TCGA-B7-5818-01:1 | Batch 57:31 |
| 5 |  | 3rd Qu.:1.0000 | D7:40 | TCGA-B7-A5TI-01:1 | Batch 95:29 |
| 6 |  | Max. :1.0000 | CD :27 | TCGA-B7-A5TJ-01:1 | Batch 242:28 |
| 7 |  |  | (Other):88 | (Other) $: 437$ | (Other) $: 187$ |

Table 4: Summary of the phenotypical information of the mRNA dataset.


Figure 2: PCA and density plot for mRNAs.

## 3 Differentially expressed miRNAs

| Analysis performed | Comparative used: CvH; method used: limma. |
| :--- | :--- |
| Number of differentially expressed miRNAs | 330 ( 150 upregulated, 180 downregulated) |
| Number of samples | 443 |
| Criteria for selecting miRNAs | adj.pval $<1$ |

Table 5: Basic statistics


Figure 3: A) Heatmap vith the top 50 most significant miRNAs (sorted by adjusted p-value). B) Volcano plot showing the selected miRNAs.

## 4 Differentially expressed mRNAs

| Analysis performed | Comparative used: CvH; method used: limma. |
| :--- | :--- |
| Number of differentially expressed mRNAs | 18565 ( 10588 upregulated, 7977 downregulated) |
| Number of samples | 443 |
| Criteria for selecting mRNAs | adj.pval $<1$ |

Table 6: Basic statistics


Figure 4: A) Heatmap vith the top 50 most significant mRNAs (sorted by adjusted p-value). B) Volcano plot showing the selected mRNAs.

## 5 Correlation \& intersection with databases

| Number of miRNAs | 330 |
| :--- | :--- |
| Number of mRNAs | 18565 |
| Total miRNA-mRNA combinations | 6126450 |
| Number of samples | 443 |

Table 7: Number of miRNAs, mRNAs and samples used for correlation.

|  | Number | \% |
| :--- | ---: | ---: |
| Total correlations | 6126450 | 100 |
| Total negative correlations | 2828695 | 46.17 |
| Total correlations p $<0.05$ | 1730533 | 28.25 |
| Total correlations p $<0.01$ | 1367300 | 22.32 |
| Total correlations adj.p $<0.05$ | 1390596 | 22.7 |
| Total correlations adj.p $<0.01$ | 1108323 | 18.09 |

Table 8: Basic statistics for correlation results. Correlation hypothesis: two.sided.


Figure 5: Density of a total of 6126450 miRNA-mRNA pairs. Dashed lines distinguish correlations whose p-value is lower than 0.05 , dotted lines for 0.01 . Black is for raw p -value and red for adjusted p -value.


Figure 6: Plot of 15 top correlations, sorted by adjusted p-value. Databases used: microCosm_v5_18, targetScan_v6.2_18 (each miRNA-mRNA pair has to appear at least 1 times).


Figure 7: Venn Diagram. Left (red): number of miRNA-mRNA pairs with ajdusted pvalue $<0.05$. Right (green): number of all the theoretical miRNA-mRNA pairs reported at least 1 times in the following databases: microCosm_v5_18, targetScan_v6.2_18. Intersection: miRNA-mRNA pairs that fulfil both conditions.


Figure 8: Circos plot for the first 45 miRNA-mRNA pairs (sorted by adjusted p-value) that have: pval-corrected $<0.05$ and appear at least 1 times in the following databases: microCosm_v5_18, targetScan_v6.2_18. Blue: miRNAs, Orange: target mRNAs

| miRNA | mRNA | cor | adj.pval | FC.miRNA | FC.mRNA | dat.sum |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| hsa-miR-141 | ZCCHC24 | -0.74 | 1.01e-72 | 1.02 | 1.03 | 1 |
| hsa-miR-141 | ZEB1 | -0.72 | $5.00 \mathrm{e}-67$ | 1.02 | 1.24 | 1 |
| hsa-miR-200c | ZCCHC24 | -0.72 | 1.46e-66 | -1.01 | 1.03 | 1 |
| hsa-miR-141 | TNS1 | -0.72 | $2.39 \mathrm{e}-65$ | 1.02 | 1.10 | 1 |
| hsa-miR-200c | DNAJB5 | -0.71 | $3.50 \mathrm{e}-65$ | -1.01 | 1.03 | 1 |
| hsa-miR-200c | KANK2 | -0.71 | $8.71 \mathrm{e}-65$ | -1.01 | 1.00 | 1 |
| hsa-miR-96 | TNS1 | -0.71 | 4.03e-64 | -1.05 | 1.10 | 1 |
| hsa-miR-200c | MYLK | -0.70 | $1.49 \mathrm{e}-62$ | -1.01 | 1.15 | 1 |
| hsa-miR-15b | LMOD1 | -0.70 | 5.93e-61 | -1.12 | -1.08 | 1 |
| hsa-miR-183 | LMOD1 | -0.70 | 5.93e-61 | 1.03 | -1.08 | 1 |
| hsa-miR-200c | ZEB1 | -0.69 | $1.32 \mathrm{e}-60$ | -1.01 | 1.24 | 2 |
| hsa-miR-96 | LMOD1 | -0.69 | $2.02 \mathrm{e}-60$ | -1.05 | -1.08 | 1 |
| hsa-miR-200b | CFL2 | -0.69 | $5.90 \mathrm{e}-60$ | -1.01 | 1.15 | 1 |
| hsa-miR-200b | DNAJB5 | -0.69 | $7.89 \mathrm{e}-60$ | -1.01 | 1.03 | 1 |
| hsa-miR-17 | FAM129A | -0.69 | $1.38 \mathrm{e}-59$ | 1.04 | 1.00 | 1 |
| hsa-miR-15b | MYLK | -0.69 | 7.07e-59 | -1.12 | 1.15 | 1 |
| hsa-miR-15b* | NEXN | -0.69 | $1.45 \mathrm{e}-58$ | -1.10 | 1.02 | 1 |
| hsa-miR-429 | CFL2 | -0.69 | $1.80 \mathrm{e}-58$ | -1.06 | 1.15 | 1 |
| hsa-miR-141 | DIXDC1 | -0.69 | $1.94 \mathrm{e}-58$ | 1.02 | 1.15 | 1 |
| hsa-miR-130b* | FHL1 | -0.68 | $1.45 \mathrm{e}-57$ | -1.04 | 1.14 | 1 |
| hsa-miR-942 | MSRB3 | -0.68 | $2.22 \mathrm{e}-57$ | -1.13 | 1.25 | 1 |
| hsa-miR-151-5p | JAM2 | -0.68 | $6.62 \mathrm{e}-57$ | 1.04 | -1.00 | 1 |
| hsa-miR-96 | MYL9 | -0.68 | $9.62 \mathrm{e}-57$ | -1.05 | -1.08 | 1 |
| hsa-miR-141 | TSHZ3 | -0.68 | $2.25 \mathrm{e}-56$ | 1.02 | 1.13 | 2 |
| hsa-miR-16 | LMOD1 | -0.68 | $2.42 \mathrm{e}-56$ | 1.03 | -1.08 | 1 |
| hsa-miR-141 | SVIL | -0.67 | $4.26 \mathrm{e}-56$ | 1.02 | 1.23 | 1 |
| hsa-miR-200a | TNS1 | -0.67 | 4.96e-56 | -1.01 | 1.10 | 1 |
| hsa-miR-200c | CFL2 | -0.67 | 5.07e-56 | -1.01 | 1.15 | 1 |
| hsa-miR-141 | MAP3K3 | -0.67 | 5.13e-56 | 1.02 | 1.06 | 1 |
| hsa-miR-141 | SYNPO2 | -0.67 | 5.34e-56 | 1.02 | 1.16 | 1 |
| hsa-miR-141 | BNC2 | -0.67 | 5.72e-56 | 1.02 | 1.31 | 1 |
| hsa-miR-15a | LMOD1 | -0.67 | $1.24 \mathrm{e}-55$ | 1.06 | -1.08 | 1 |
| hsa-miR-106b | TNS1 | -0.67 | $1.34 \mathrm{e}-55$ | -1.04 | 1.10 | 1 |
| hsa-miR-429 | MYLK | -0.67 | $1.57 \mathrm{e}-55$ | -1.06 | 1.15 | 1 |
| hsa-miR-141 | NECAB1 | -0.67 | $1.87 \mathrm{e}-55$ | 1.02 | 1.10 | 1 |
| hsa-miR-200b* | GPRASP1 | -0.67 | 3.08e-55 | -1.11 | 1.13 | 1 |
| hsa-miR-429 | DNAJB5 | -0.67 | $3.59 \mathrm{e}-55$ | -1.06 | 1.03 | 1 |
| hsa-miR-200c | POU6F1 | -0.67 | $1.09 \mathrm{e}-54$ | -1.01 | 1.07 | 1 |
| hsa-miR-200a | SVIL | -0.67 | $1.14 \mathrm{e}-54$ | -1.01 | 1.23 | 1 |
| hsa-miR-148b | CNN1 | -0.67 | 1.73e-54 | -1.05 | -1.15 | 1 |
| hsa-miR-200b | NDN | -0.67 | $2.37 \mathrm{e}-54$ | -1.01 | -1.04 | 2 |
| hsa-miR-15b | PRELP | -0.67 | $3.45 \mathrm{e}-54$ | -1.12 | 1.11 | 1 |
| hsa-miR-576-5p | NEGR1 | -0.67 | $3.77 \mathrm{e}-54$ | 1.04 | 1.06 | 1 |
| hsa-miR-106b | CNN1 | -0.66 | $5.20 \mathrm{e}-54$ | -1.04 | -1.15 | 2 |
| hsa-miR-16 | MYLK | -0.66 | $5.89 \mathrm{e}-54$ | 1.03 | 1.15 | 1 |

Table 9: Top 45 miRNA-mRNA pairs(sorted by adjusted p-value) that have: pvalcorrected $<0.05$ and appear at least 1 times in the following databases: microCosm_v5_18, targetScan_v6.2_18.

## 6 Functional analysis

### 6.1 Network analysis



Figure 9: Network for all the miRNA-mRNA pairs that have: pval-corrected $<0.05$ and appear at least 1 times in the following databases: microCosm_v5_18, targetScan_v6.2_18. Circles represent the miRNAs, and squares the miRNA. Red fill means upregulated miRNAs/mRNAs, while green fill means downregulated mRNA/mRNAs in comparative CvH ; lines indicate the miRNA-mRNA pairs, red line means positive score and green line means negative score.


Figure 10: Barplot for miRNAs, pval-corrected<0.05 and Targets=microCosm_v5_18, targetScan_v6.2_18(minimum coincidences between databases:1). Red line (and right axis) represents the percentage of deregulated mRNAs that are targeted by the miRNAs.


Figure 11: Pie chart representing the number of miRNAs targeting the mRNAs, pvalcorrected $<0.05$ and Targets=microCosm_v5_18, targetScan_v6.2_18(minimum coincidences between databases:1).

| miRNA | \#targets | cum. \% | targets (top 20) |
| :--- | ---: | ---: | :--- | :--- |
| hsa-miR-29a | 798 | 4.3 | FEM1B, LAMC1, STRN3, FAM168B, PRKAB2, |
|  |  |  | CBX1, ITGB1, KPNA4, TMTC3, ARNT, RBFOX2, |
|  |  |  | ENAH, SESTD1, WDR26, C5orf24, KDELC1, ELF2, |
|  |  | DIP2C, BTBD7, JMY |  |
| hsa-miR-29c | 784 | 5.93 | COL11A1, FBXO45, PLEKHA8, KDELC1, CCNJ, |
|  |  |  | CSE1L, LRRC58, MTHFD1L, CALU, KPNA4, |
|  |  |  | USP31, CBX1, BMP1, TRIM37, XPO5, ABCE1, |
|  |  | RIOK1, ZC3HAV1L, NKRF, SYNCRIP |  |

Table 10: Top 10 miRNA with more targets (each miRNA-mRNA pair has pvalcorrected $<0.05$ and appears at least 1 times in the following databases: microCosm_v5_18, targetScan_v6.2_18). MiRNAs in red are upregulated in CvH, miRNAs in green are downregulated in CvH .

| mRNA | \#miRNAs | miRNAs (top 20) |
| :---: | :---: | :---: |
| FOXP2 | 74 | hsa-miR-15b, hsa-miR-625, hsa-miR-141, hsa-miR-16, hsa-miR-130b*, hsa-miR-590-3p, hsa-miR-7-1*, hsa-miR-576-5p, hsa-miR-15a, hsa-miR-222, hsa-miR-502-3p, hsa-miR-96, hsa-miR-200a, hsa-miR-590-5p, hsa-let-7d, hsa-miR-671-5p, hsa-miR-7, hsa-miR-34a, hsa-miR-197, hsa-miR-182 |
| TNRC6B | 67 | hsa-miR-130b, hsa-miR-26b, hsa-miR-144, hsa-miR-32, hsa-miR-590-3p, hsa-miR-106b, hsa-miR-16, hsa-miR-7, hsa-miR-3613-5p, hsa-miR-141, hsa-miR-424, hsa-miR-183, hsa-miR-29b, hsa-miR-29a, hsa-miR-590-5p, hsa-miR-455-3p, hsa-miR-3614-5p, hsa-miR-15b, hsa-miR-330-3p, hsa-miR-18a |
| QKI | 65 | hsa-miR-429, hsa-miR-141, hsa-miR-200a, hsa-miR-200b, hsa-miR-200c, hsa-miR-96, hsa-miR-532-3p, hsa-miR-194, hsa-miR-130b, hsa-miR-19b, hsa-miR-33a, hsa-miR-92a, hsa-miR-106b, hsa-miR-345, hsa-miR-17, hsa-miR-93*, hsa-miR-500b, hsa-miR-148a, hsa-miR-577, hsa-miR-19a |
| RORA | 65 | hsa-miR-141, hsa-miR-106b, hsa-miR-18a, hsa-miR-17, hsa-miR-200a, hsa-miR-15b, hsa-miR-33a, hsa-miR-3613-5p, hsa-miR-19b, hsa-miR-92a, hsa-miR-576-5p, hsa-miR-590-3p, hsa-miR-502-3p, hsa-miR-671-5p, hsa-miR652, hsa-miR-15a, hsa-miR-183, hsa-miR-19a, hsa-miR-93, hsa-miR-501-3p |
| BNC2 | 60 | hsa-miR-141, hsa-miR-200c, hsa-miR-130b, hsa-miR-106b, hsa-miR-429, hsa-miR-96, hsa-miR-200a, hsa-miR-592, hsa-miR-200b, hsa-miR-7, hsa-miR-17, hsa-miR-19b, hsa-miR-183, hsa-miR-200c*, hsa-miR-191, hsa-miR-577, hsa-miR-590-5p, hsa-miR-7-1*, hsa-miR-182, hsa-miR-19a |
| AFF4 | 57 | hsa-miR-30b, hsa-miR-374b, hsa-miR-106b, hsa-miR-92a, hsa-miR-29a, hsa-miR-429, hsa-miR-32, hsa-miR-197, hsa-miR-590-3p, hsa-miR-4555p, hsa-miR-502-3p, hsa-miR-15b, hsa-miR-200c, hsa-miR-17, hsa-miR-25, hsa-miR-505, hsa-miR-374a, hsa-miR-660, hsa-miR-15a, hsa-miR-331-5p |
| LPP | 57 | hsa-miR-590-3p, hsa-miR-16, hsa-miR-141, hsa-miR-96, hsa-miR-942, hsa-miR-32, hsa-miR-183, hsa-miR-7, hsa-miR-1266, hsa-miR-15b, hsa-miR182, hsa-miR-424, hsa-miR-1976, hsa-miR-576-5p, hsa-miR-19b, hsa-miR18a, hsa-miR-425, hsa-miR-330-5p, hsa-miR-577, hsa-miR-200a |
| RUNX1T1 | 57 | hsa-miR-15b, hsa-miR-130b, hsa-miR-16, hsa-miR-200c, hsa-miR-33a, hsa-miR-429, hsa-miR-148b, hsa-miR-455-3p, hsa-miR-32, hsa-miR-200b, hsa-miR-19b, hsa-miR-15a, hsa-miR-92a, hsa-miR-19a, hsa-let-7d, hsa-miR192, hsa-miR-25, hsa-miR-107, hsa-miR-29b, hsa-miR-186 |
| CPEB4 | 55 | hsa-miR-92a, hsa-miR-942, hsa-miR-19b, hsa-miR-130b, hsa-miR-197, hsa-miR-19a, hsa-miR-500b, hsa-miR-27a, hsa-miR-362-5p, hsa-miR-25, hsa-miR-96, hsa-miR-550a, hsa-miR-590-3p, hsa-miR-3647-3p, hsa-miR-128, hsa-let-7d, hsa-miR-26b, hsa-miR-141, hsa-miR-374b, hsa-miR-192 |
| NFIA | 55 | hsa-miR-130b, hsa-miR-590-5p, hsa-miR-141, hsa-miR-32, hsa-miR-942, hsa-miR-93*, hsa-miR-191, hsa-miR-7, hsa-miR-33a, hsa-miR-454, hsa-miR-25, hsa-miR-200c, hsa-miR-429, hsa-miR-19a, hsa-miR-19b, hsa-miR301a, hsa-miR-330-3p, hsa-miR-577, hsa-miR-200a, hsa-miR-92a |

Table 11: Top 10 mRNA with more miRNAs targeting them (each miRNA-mRNA pair has pval-corrected $<0.05$ and appears at least 1 times in the following databases: microCosm_v5_18, targetScan_v6.2_18). MRNAs in red are upregulated in CvH, mRNAs in green are downregulated in CvH .

### 6.2 GO analysis

| GOBPID | Term | Count | Size | ExpCount | OddsRatio | fdr | Pvalue |
| :--- | :--- | ---: | ---: | ---: | ---: | ---: | ---: |
| GO:0009987 | cellular process | 8586 | 12932 | 8354.90 | 1.91 | $6.41 \mathrm{e}-31$ | $5.53 \mathrm{e}-35$ |
| GO:0006464 | cellular protein modifi- <br> cation process | 1931 | 2583 | 1668.78 | 1.78 | $1.51 \mathrm{e}-30$ | $3.90 \mathrm{e}-34$ |
| GO:0036211 | protein modification <br> process | 1931 | 2583 | 1668.78 | 1.78 | $1.51 \mathrm{e}-30$ | $3.90 \mathrm{e}-34$ |
| GO:0016043 | cellular component or- <br> ganization | 3304 | 4619 | 2984.17 | 1.58 | $5.28 \mathrm{e}-30$ | $1.82 \mathrm{e}-33$ |
| GO:0071840 | cellular component or- <br> ganization or biogenesis | 3365 | 4715 | 3046.19 | 1.57 | $1.57 \mathrm{e}-29$ | $6.78 \mathrm{e}-33$ |
| GO:0043412 | macromolecule modifi- <br> cation | 1994 | 2685 | 1734.68 | 1.74 | $4.11 \mathrm{e}-29$ | $2.13 \mathrm{e}-32$ |
| GO:0044267 | cellular <br> metabolic process | 2456 | 3370 | 2177.24 | 1.64 | $3.37 \mathrm{e}-28$ | $2.03 \mathrm{e}-31$ |
| GO:0006793 | phosphorus metabolic <br> process | 2061 | 2809 | 1814.79 | 1.65 | $3.34 \mathrm{e}-25$ | $2.51 \mathrm{e}-28$ |
| GO:0050794 | regulation of cellular <br> process | 5447 | 7941 | 5130.40 | 1.46 | $3.34 \mathrm{e}-25$ | $2.59 \mathrm{e}-28$ |
| GO:0044237 | cellular metabolic pro- <br> cess | 6031 | 8855 | 5720.90 | 1.47 | $4.77 \mathrm{e}-25$ | $4.12 \mathrm{e}-28$ |

Table 12: Biological Process . Options used: mRNAs that are present in a mRNAmRNA pair that has adjusted-pval cutoff $<0.05$; that also appears at least 1 times (databases: microCosm_v5_18, targetScan_v6.2_18); organism: human.

| GOCCID | Term | Count | Size | ExpCount | OddsRatio | fdr | Pvalue |
| ---: | :--- | ---: | ---: | ---: | ---: | ---: | ---: | ---: |
| GO:0044424 | intracellular part | 8246 | 12258 | 7540.33 | 2.46 | $2.62 \mathrm{e}-137$ | $1.95 \mathrm{e}-140$ |
| GO:0005622 | intracellular | 8322 | 12396 | 7625.22 | 2.47 | $7.87 \mathrm{e}-137$ | $1.17 \mathrm{e}-139$ |
| GO:0005737 | cytoplasm | 6465 | 9342 | 5746.60 | 2.10 | $7.01 \mathrm{e}-115$ | $1.56 \mathrm{e}-117$ |
| GO:0043226 | organelle | 7566 | 11215 | 6898.74 | 2.13 | $1.04 \mathrm{e}-109$ | $3.10 \mathrm{e}-112$ |
| GO:0043227 | membrane-bounded or- | 7078 | 10399 | 6396.79 | 2.08 | $6.74 \mathrm{e}-108$ | $2.50 \mathrm{e}-110$ |
|  | ganelle |  |  |  |  |  |  |
| GO:0043229 | intracellular organelle | 7191 | 10617 | 6530.89 | 2.05 | $9.60 \mathrm{e}-103$ | $4.28 \mathrm{e}-105$ |
| GO:0043231 | intracellular membrane- | 6546 | 9551 | 5875.16 | 2.00 | $5.33 \mathrm{e}-101$ | $2.78 \mathrm{e}-103$ |
|  | bounded organelle |  |  |  |  |  |  |
| GO:0044444 | cytoplasmic part | 4827 | 6864 | 4222.29 | 1.91 | $2.79 \mathrm{e}-84$ | $1.66 \mathrm{e}-86$ |
| GO:0044446 | intracellular organelle | 4392 | 6274 | 3859.36 | 1.80 | $1.26 \mathrm{e}-67$ | $8.44 \mathrm{e}-70$ |
|  | part | 4502 | 6449 | 3967.01 | 1.80 | $1.78 \mathrm{e}-67$ | $1.33 \mathrm{e}-69$ |

Table 13: Cellular Component . Options used: mRNAs that are present in a mRNAmRNA pair that has adjusted-pval cutoff $<0.05$; that also appears at least 1 times (databases: microCosm_v5_18, targetScan_v6.2_18); organism: human.

| GOMFID | Term | Count | Size | ExpCount | OddsRatio | fdr | Pvalue |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| GO:0005515 | protein binding | 5593 | 7904 | 4974.68 | 2.03 | 2.25e-93 | 6.53e-97 |
| GO:0005488 | binding | 8058 | 12075 | 7599.85 | 2.18 | 5.07e-77 | $2.95 \mathrm{e}-80$ |
| GO:0043167 | ion binding | 3986 | 5793 | 3646.04 | 1.51 | $1.80 \mathrm{e}-29$ | 1.57e-32 |
| GO:0043168 | anion binding | 1821 | 2493 | 1569.06 | 1.74 | $1.28 \mathrm{e}-28$ | $1.49 \mathrm{e}-31$ |
| GO:1901265 | nucleoside phosphate binding | 1660 | 2255 | 1419.27 | 1.78 | $1.52 \mathrm{e}-28$ | $2.21 \mathrm{e}-31$ |
| GO:0000166 | nucleotide binding | 1659 | 2254 | 1418.64 | 1.78 | $1.53 \mathrm{e}-28$ | $2.67 \mathrm{e}-31$ |
| GO:0017076 | purine nucleotide binding | 1348 | 1819 | 1144.86 | 1.80 | $1.77 \mathrm{e}-24$ | $3.61 \mathrm{e}-27$ |
| GO:0097367 | carbohydrate derivative binding | 1548 | 2115 | 1331.15 | 1.73 | $1.87 \mathrm{e}-24$ | $4.34 \mathrm{e}-27$ |
| GO:0003824 | catalytic activity | 3571 | 5197 | 3270.93 | 1.47 | 2.28e-24 | $5.96 \mathrm{e}-27$ |
| GO:0032555 | purine ribonucleotide binding | 1334 | 1800 | 1132.90 | 1.80 | $2.36 \mathrm{e}-24$ | $6.84 \mathrm{e}-27$ |

Table 14: Molecular Function . Options used: mRNAs that are present in a mRNAmRNA pair that has adjusted-pval cutoff $<0.05$; that also appears at least 1 times (databases: microCosm_v5_18, targetScan_v6.2_18); organism: human.

| KEGGID | Term | Count | Size | ExpCount | OddsRatio | fdr | Pvalue |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| 05200 | Pathways in cancer | 253 | 314 | 201.06 | 2.43 | 6.42e-09 | 3.88e-11 |
| 04510 | Focal adhesion | 163 | 192 | 122.94 | 3.26 | 6.42e-09 | 5.65e-11 |
| 04722 | Neurotrophin signaling pathway | 105 | 125 | 80.04 | 3.01 | $3.89 \mathrm{e}-05$ | 5.14e-07 |
| 04810 | Regulation of actin cytoskeleton | 161 | 203 | 129.99 | 2.21 | 5.85e-05 | 1.14e-06 |
| 04141 | Protein processing in endoplasmic reticulum | 132 | 163 | 104.37 | 2.44 | 5.85e-05 | 1.29e-06 |
| 05100 | Bacterial invasion of epithelial cells | 61 | 69 | 44.18 | 4.34 | 1.58e-04 | 4.18e-06 |
| 04666 | Fc gamma R-mediated phagocytosis | 78 | 92 | 58.91 | 3.18 | 2.45e-04 | 7.57e-06 |
| 05222 | Small cell lung cancer | 71 | 83 | 53.15 | 3.37 | 2.51e-04 | 1.01e-05 |
| 05215 | Prostate cancer | 74 | 87 | 55.71 | 3.24 | 2.51e-04 | 1.02e-05 |
| 04360 | Axon guidance | 102 | 125 | 80.04 | 2.53 | $2.51 \mathrm{e}-04$ | 1.11e-05 |

Table 15: Kegg Pathways. Options used: mRNAs that are present in a mRNA-mRNA pair that has adjusted-pval cutoff $<0.05$; that also appears at least 1 times (databases: microCosm_v5_18, targetScan_v6.2_18); organism: human.

