

Supplementary material

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Supplementary Table 1. Differentially expressed genes in two statistical tests

| | Log ₂ (FC) ^{1*} | Odds ratio | P value in | P value in |
|------------|-------------------------------------|------------|------------|--------------|
| | | | logistic | Kolmogorov- |
| | | | regression | Smirnov test |
| RPL15 | 0.331 | 1.479 | 0.022 | 0.005 |
| BACE1-AS | 0.271 | 1.384 | 0.034 | 0.013 |
| CEPT1 | 0.105 | 1.416 | 0.022 | 0.030 |
| EIF4G1 | 0.573 | 1.702 | 0.015 | 0.044 |
| TMEM91 | 0.357 | 1.419 | 0.027 | 0.044 |
| TBCK | 0.519 | 1.344 | 0.036 | 0.044 |
| ARG2 | 0.135 | 2.137 | 0.161 | 0.010 |
| DIS3 | 0.280 | 1.904 | 0.063 | 0.020 |
| TMEM 129 | 0.181 | 2.368 | 0.101 | 0.028 |
| PRPF40B | 0.270 | 2.763 | 0.115 | 0.030 |
| PCCB | 0.225 | 1.279 | 0.105 | 0.030 |
| PASK | 0.278 | 2.054 | 0.214 | 0.030 |
| KCNQ5 | 0.316 | 1.428 | 0.093 | 0.030 |
| ATP1A1-AS1 | 0.053 | 2.798 | 0.230 | 0.040 |
| FAM120AOS | 0.238 | 1.182 | 0.157 | 0.044 |

Abbreviation: $Log_2(FC) = log_2(fold-change)$

^{*} Value >0 indicates higher mean expression in the severe group

Supplementary Table 2. Differentially expressed genes in the 46-65 age group*

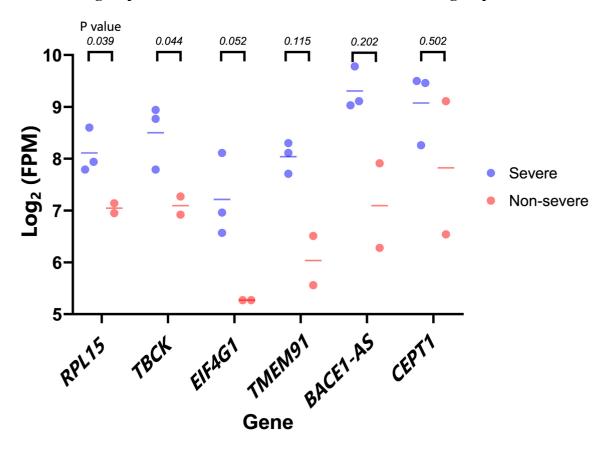
| | Severe group log2(FPM) values | | Non-severe group log2(FPM) values | | Log ₂ (FC) | P value | |
|----------|----------------------------------|--------|-----------------------------------|-----------|-----------------------|---------|------------|
| | Case 1 | Case 2 | Case 3 | Control 1 | Control 2 | | in t tests |
| RPL15 | 7.79 | 8.6 | 7.94 | 6.95 | 7.14 | 0.20 | 0.039 |
| TBCK | 7.79 | 8.77 | 8.94 | 7.27 | 6.92 | 0.26 | 0.044 |
| EIF4G1 | 6.57 | 6.96 | 8.11 | 5.27 | 5.27^{\dagger} | 0.45 | 0.052 |
| TMEM91 | 8.3 | 7.71 | 8.11 | 5.56 | 6.51 | 0.41 | 0.115 |
| BACE1-AS | 9.78 | 9.03 | 9.11 | 6.28 | 7.91 | 0.39 | 0.202 |
| CEPT1 | 9.5 | 9.46 | 8.26 | 6.54 | 9.11 | 0.21 | 0.502 |

Abbreviations: $Log_2(FC) = log_2(fold\text{-change}); log_2(FPM) = log_2(fragments per million reads mapped)$

^{*}Among the 24 patients, three did not pass the quality control requirement for RNA sequencing data. One patient without age information was excluded. Among the remaining 23 patients, there were zero severe cases in the 0-18 age group, zero severe cases in the >65 age group, and one severe case in the 19-45 age group; these values were insufficient for statistical analyses. Among the remaining six patients in the 46-65 age group, one control patient (in the non-severe group) had zero counts for these genes and was excluded. Thus, t tests on log_2 values were performed between the three patients with severe disease and the two patients with non-severe disease

[†] Missing value imputed using the mean expression level for this gene in the 16-65 age group

Supplementary Figure. Differentially expressed genes in the 46-65 age group. Horizontal lines indicate group mean. P values in t tests are shown above the groups



References

1. Mortazavi A, Williams BA, McCue K, Schaeffer L, Wold B. Mapping and quantifying mammalian transcriptomes by RNA-Seq. Nat Methods 2008;5:621-8.