|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
|  | **Paired t-test**  **(Follow-up – Baseline)** | | **Correlation test** | | |
| **Metabolites** | **t-statistic** | ***p*-value** | **rPearson** | | ***p*-value** |
| 1,5-anhydroglucitol | -3.88 ↘ | 1.13×10-4 | **0.73** | **7.29×10-139** | |
| 1-linoleoyl-GPC | -2.62 ↘ | 8.89×10-3 | 0.40 | 2.46×10-33 | |
| 1-palmitoylglycerol | 5.12 ↗ | 3.81×10-7 | 0.45 | 1.55×10-42 | |
| Cotinine | -5.32 ↘ | 1.32×10-7 | 0.63 | 1.05×10-94 | |
| γ-glutamylphenylalanine | 2.20 ↗ | 0.03 | 0.40 | 8.24×10-34 | |
| Glucose | 6.72 ↗ | 3.34×10-11 | 0.40 | 5.30×10-33 | |
| Isoleucine | 6.94 ↗ | 7.89×10-12 | 0.56 | 2.78×10-70 | |
| Mannose | 6.59 ↗ | 7.70×10-11 | 0.50 | 2.59×10-54 | |
| Pro-hydroxy-pro | 6.78 ↗ | 2.29×10-11 | 0.39 | 4.75×10-31 | |
| Fructose | -0.05 | 0.96 | 0.05 | 0.15 | |
| γ-glutamyltyrosine | 9.54 ↗ | 1.47×10-20 | 0.46 | 6.57×10-45 | |
| Isovalerylcarnitine | 4.33 ↗ | 1.65×10-5 | 0.57 | 2.13×10-72 | |
| Phenylalanine | -0,51 | 0.61 | 0.43 | 1.13×10-38 | |
| Piperine | 1.40 | 0.16 | 0.23 | 2.26×10-11 | |
| Serine | -4.12 ↘ | 4.25×10-5 | 0.26 | 4×10-14 | |
| Tyrosine | 6.77 ↗ | 2.38×10-11 | 0.53 | 6.70×10-61 | |
| 1-stearoyl-GPI | 1.15 ↗ | 0.25 | 0.08 | 0.02 | |
| 3-hydroxyisobutyrate | 2.49 ↗ | 0.01 | 0.42 | 2.56×10-37 | |
| Dehydroisoandrosterone sulfate | -19.37 ↘ | 2.66×10-69 | **0.84** | **5.44×10-221** | |
| γ-glutamylvaline | -7.73 ↘ | 3×10-14 | 0.21 | 1.44×10-9 | |
| Glycine | -7.75 ↘ | 2.76×10-14 | 0.55 | 4.24×10-66 | |
| Palmitoyl sphingomyelin | 0.25 | 0.08 | 0.36 | 1.40×10-26 | |
| Stearoylcarnitine | -0.54 | 0.59 | 0.18 | 1.09×10-7 | |
| Urea | 7.92 ↗ | 7.55×10-15 | 0.50 | 2.29×10-53 | |

**Supplementary Table 3**. T-statistics and *p*-values compare average levels in metabolites contributing to MRS1 and MRS2, at baseline and at 9 years. Pearson correlation coefficients (*p*-values) are also provided. ↘ and ↗ mean that metabolite levels have decreased or increased respectively nine years after inclusion. The D.E.S.I.R. study.