

SUPPLEMENTARY MATERIALS:

New insights on the PBMCs phospholipidome in obesity demonstrate modulations associated with insulin resistance and glycemic status

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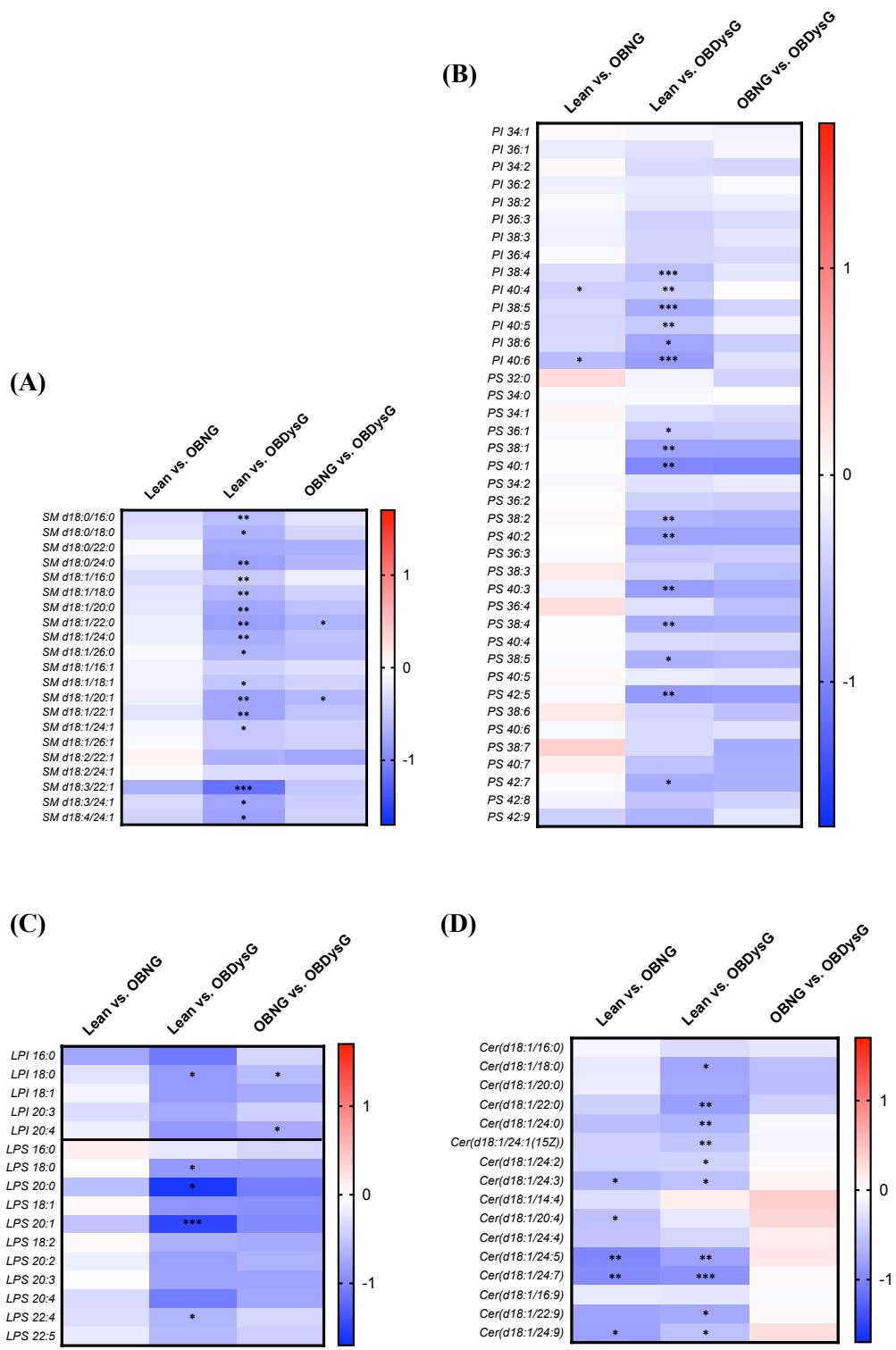


Figure S1. Fold change (Log2FC) of all (A) SM, (B) PI/PS, (C) Lyso-PI/-PS and (D) Cer lipid species detected in PBMCs from lean compared to OBNG or OBDysG patients and from OBNG vs. OBDysG individuals. Mean comparison * p≤0.05; **p ≤ 0.01; ***p ≤ 0.001. Kruskal-Wallis followed by Dunn post hoc test was performed on data.

Table S1. Participants blood cells composition. Data are mean \pm SD. Kruskal-Wallis followed by Dunn post hoc test was performed on data. OBNG or OBIG vs. Lean * $p \leq 0.05$; ** $p \leq 0.01$; *** $p \leq 0.001$. OBIG vs. OBNG † $p \leq 0.05$; ‡‡ $p \leq 0.01$; ‡‡‡ $p \leq 0.001$.

| | Overall p-value | Lean | OBNG | OBDysG |
|---|-----------------|-------------------|--------------------|-------------------|
| Platelets ($10^3/\text{mm}^3$) | 0.238 | 244.6 ± 37.7 | 293.7 ± 104.0 | 255.6 ± 50.0 |
| Leucocytes ($10^3/\text{mm}^3$) | 0.004 | 6.1 ± 1.2 | 7.7 ± 2.0 * | 8.0 ± 1.8 ** |
| Neutrophils (%) | 0.375 | 56.8 ± 9.2 | 56.8 ± 6.0 | 59.2 ± 7.1 |
| Lymphocytes (%) | 0.22 | 32.5 ± 7.2 | 33.2 ± 6.4 | 29.5 ± 6.6 |
| Monocytes (%) | 0.275 | 7.94 ± 1.39 | 7.07 ± 1.79 | 7.59 ± 2.14 |
| Eosinophils (%) | 0.216 | 2.27 ± 1.65 | 2.27 ± 1.40 | 2.82 ± 1.38 |
| Basophiles (%) | 0.869 | 0.48 ± 0.22 | 0.54 ± 0.36 | 0.52 ± 0.22 |
| Neutrophils absolute ($10^3/\text{mm}^3$) | 0.024 | 3.53 ± 1.19 | 4.36 ± 1.32 | 4.79 ± 1.49 * |
| Lymphocytes absolute ($10^3/\text{mm}^3$) | 0.012 | 1.92 ± 0.36 | 2.55 ± 0.76 ** | 2.32 ± 0.65 |
| Monocytes absolute ($10^3/\text{mm}^3$) | 0.096 | 0.47 ± 0.09 | 0.54 ± 0.19 | 0.60 ± 0.18 |
| Eosinophils absolute ($10^3/\text{mm}^3$) | 0.017 | 0.13 ± 0.09 | 0.17 ± 0.12 | 0.21 ± 0.12 * |
| Basophiles absolute ($10^3/\text{mm}^3$) | 0.102 | 0.029 ± 0.013 | 0.040 ± 0.027 | 0.255 ± 1.035 |

Table S2. Individual PBMCs lipid species. Data are mean ± SD.

| Lipid species (nmol/mg DNA) | Lean | OBNG | OBIG |
|--------------------------------|------------------|-----------------|-----------------|
| PC 28:0 | 0.143 ± 0.062 | 0.13 ± 0.073 | 0.08 ± 0.057 |
| PC 30:0 | 2.882 ± 0.886 | 2.299 ± 0.931 | 1.845 ± 0.742 |
| PC 32:0 | 61.659 ± 13.396 | 50.188 ± 13.016 | 46.932 ± 14.065 |
| PC 34:0 | 13.566 ± 3.539 | 10.416 ± 3.361 | 8.831 ± 3.513 |
| PC 36:0 | 2.133 ± 0.765 | 1.667 ± 0.707 | 1.277 ± 0.724 |
| PC 38:0 | 1.882 ± 0.422 | 1.476 ± 0.512 | 1.437 ± 0.621 |
| PC 40:0 | 0.307 ± 0.152 | 0.232 ± 0.153 | 0.163 ± 0.116 |
| PC 32:1 | 6.713 ± 1.81 | 6.242 ± 2.272 | 4.921 ± 1.835 |
| PC 34:1 | 143.536 ± 41.405 | 118.489 ± 40.52 | 95.26 ± 35.32 |
| PC 36:1 | 49.229 ± 13.757 | 39.824 ± 14.686 | 30.659 ± 13.046 |
| PC 40:1 | 0.421 ± 0.118 | 0.346 ± 0.13 | 0.263 ± 0.103 |
| PC 32:2 | 0.655 ± 0.213 | 0.522 ± 0.283 | 0.35 ± 0.161 |
| PC 34:2 | 50.128 ± 13.158 | 37.071 ± 12.733 | 29.893 ± 12.119 |
| PC 36:2 | 62.639 ± 11.081 | 49.946 ± 12.957 | 42.966 ± 14.997 |
| PC 38:2 | 4.763 ± 1.25 | 3.321 ± 1.348 | 2.564 ± 1.272 |
| PC 40:2 | 0.459 ± 0.148 | 0.376 ± 0.153 | 0.285 ± 0.119 |
| PC 42:2 | 0.345 ± 0.099 | 0.29 ± 0.113 | 0.225 ± 0.088 |
| PC 44:2 | 0.354 ± 0.226 | 0.282 ± 0.111 | 0.222 ± 0.122 |
| PC 32:3 | 0.182 ± 0.079 | 0.138 ± 0.056 | 0.111 ± 0.058 |
| PC 34:3 | 2.515 ± 0.618 | 2.097 ± 0.863 | 1.511 ± 0.637 |
| PC 36:3 | 32.44 ± 5.272 | 26.618 ± 7.242 | 21.573 ± 7.474 |
| PC 38:3 | 13.762 ± 2.533 | 12.099 ± 3.798 | 9.096 ± 3.272 |
| PC 40:3 | 0.789 ± 0.214 | 0.702 ± 0.306 | 0.498 ± 0.189 |
| PC 42:3 | 0.258 ± 0.09 | 0.237 ± 0.074 | 0.183 ± 0.079 |
| PC 44:3 | 0.28 ± 0.187 | 0.227 ± 0.095 | 0.174 ± 0.105 |
| PC 34:4 | 0.824 ± 0.282 | 0.663 ± 0.312 | 0.474 ± 0.243 |
| PC 36:4 | 65.946 ± 15.286 | 54.066 ± 16.761 | 46.74 ± 15.803 |
| PC 38:4 | 77.49 ± 17.416 | 64.194 ± 17.667 | 56.402 ± 20.47 |
| PC 40:4 | 6.039 ± 1.717 | 5.151 ± 1.923 | 3.965 ± 1.699 |
| PC 42:4 | 0.503 ± 0.152 | 0.433 ± 0.169 | 0.318 ± 0.14 |
| PC 44:4 | 0.2 ± 0.152 | 0.179 ± 0.08 | 0.128 ± 0.069 |
| PC 36:5 | 3.641 ± 0.935 | 3.088 ± 1.121 | 2.501 ± 1.045 |
| PC 38:5 | 36.565 ± 5.891 | 30.847 ± 8.617 | 26.964 ± 9.185 |
| PC 40:5 | 10.59 ± 2.413 | 9.141 ± 3.082 | 7.745 ± 2.921 |
| PC 42:5 | 0.869 ± 0.235 | 0.741 ± 0.304 | 0.583 ± 0.243 |
| PC 44:5 | 0.399 ± 0.219 | 0.317 ± 0.128 | 0.243 ± 0.133 |
| PC 36:6 | 0.39 ± 0.175 | 0.334 ± 0.16 | 0.25 ± 0.135 |
| PC 38:6 | 11.553 ± 2.458 | 8.6 ± 2.65 | 7.462 ± 2.647 |

| Lipid species (nmol/mg DNA) | Lean | OBNG | OBIG |
|--------------------------------|--------------------------|-----------------------|--------------------------|
| PC 40:6 | 9.497 ± 1.969 | 7.518 ± 2.291 | 6.579 ± 2.247 |
| PC 42:6 | 1.032 ± 0.249 | 0.86 ± 0.334 | 0.673 ± 0.262 |
| PC 44:6 | 0.469 ± 0.218 | 0.362 ± 0.184 | 0.267 ± 0.157 |
| PC 38:7 | 1.002 ± 0.299 | 0.842 ± 0.311 | 0.642 ± 0.276 |
| PC 40:7 | 4.903 ± 1.337 | 4.082 ± 1.567 | 3.171 ± 1.295 |
| PC 42:7 | 1.167 ± 0.291 | 0.985 ± 0.374 | 0.771 ± 0.279 |
| PC 44:7 | 0.319 ± 0.151 | 0.256 ± 0.113 | 0.198 ± 0.1 |
| PC 40:8 | 2.403 ± 0.7 | 2.023 ± 0.863 | 1.56 ± 0.717 |
| PC 42:8 | 1.336 ± 0.388 | 1.144 ± 0.514 | 0.876 ± 0.369 |
| PC 44:8 | 0.437 ± 0.166 | 0.418 ± 0.153 | 0.346 ± 0.149 |
| PC 42:9 | 1.031 ± 0.306 | 0.964 ± 0.397 | 0.753 ± 0.379 |
| PC 44:9 | 0.367 ± 0.123 | 0.338 ± 0.122 | 0.255 ± 0.111 |
| PC 42:10 | 0.953 ± 0.281 | 0.854 ± 0.323 | 0.7 ± 0.374 |
| PC 44:10 | 0.47 ± 0.147 | 0.396 ± 0.151 | 0.311 ± 0.119 |
| PC 42:11 | 0.898 ± 0.256 | 0.812 ± 0.29 | 0.658 ± 0.318 |
| PC 44:11 | 0.748 ± 0.259 | 0.681 ± 0.238 | 0.557 ± 0.28 |
| PC 42:12 | 1.36 ± 0.431 | 1.179 ± 0.401 | 1.028 ± 0.469 |
| PC - total | 695.439 ± 139.616 | 566.7 ± 166.05 | 474.443 ± 161.902 |
| SM d18:0/16:0 | 1.887 ± 0.539 | 1.526 ± 0.546 | 1.295 ± 0.402 |
| SM d18:0/18:0 | 0.539 ± 0.21 | 0.447 ± 0.221 | 0.347 ± 0.164 |
| SM d18:0/22:0 | 2.093 ± 1.181 | 2.002 ± 1.342 | 1.267 ± 0.764 |
| SM d18:0/24:0 | 1.051 ± 0.442 | 0.938 ± 0.479 | 0.616 ± 0.307 |
| SM d18:1/16:0 | 29.992 ± 6.904 | 24.492 ± 6.379 | 22.086 ± 7.412 |
| SM d18:1/18:0 | 2.391 ± 0.766 | 2.041 ± 0.715 | 1.565 ± 0.707 |
| SM d18:1/20:0 | 5.047 ± 1.881 | 4.342 ± 2.236 | 3.027 ± 1.517 |
| SM d18:1/22:0 | 15.339 ± 6.138 | 13.837 ± 7.397 | 8.886 ± 4.51 |
| SM d18:1/24:0 | 11.271 ± 3.869 | 10.233 ± 4.429 | 7.087 ± 3.257 |
| SM d18:1/26:0 | 0.243 ± 0.088 | 0.233 ± 0.109 | 0.158 ± 0.075 |
| SM d18:1/16:1 | 1.506 ± 0.479 | 1.386 ± 0.471 | 1.154 ± 0.458 |
| SM d18:1/18:1 | 0.636 ± 0.209 | 0.587 ± 0.199 | 0.453 ± 0.206 |
| SM d18:1/20:1 | 0.683 ± 0.229 | 0.614 ± 0.254 | 0.41 ± 0.198 |
| SM d18:1/22:1 | 6.084 ± 1.983 | 5.168 ± 2.458 | 3.63 ± 1.628 |
| SM d18:1/24:1 | 23.002 ± 6.985 | 21.629 ± 8.308 | 16.664 ± 6.955 |
| SM d18:1/26:1 | 0.514 ± 0.162 | 0.492 ± 0.195 | 0.374 ± 0.153 |
| SM d18:2/22:1 | 1.533 ± 0.989 | 1.65 ± 1.245 | 0.981 ± 0.633 |
| SM d18:2/24:1 | 5.907 ± 1.733 | 5.951 ± 2.158 | 4.785 ± 1.965 |
| SM d18:3/22:1 | 1.08 ± 0.509 | 0.687 ± 0.424 | 0.492 ± 0.288 |
| SM d18:3/24:1 | 1.891 ± 0.725 | 1.524 ± 0.76 | 1.142 ± 0.721 |
| SM d18:4/24:1 | 0.744 ± 0.346 | 0.571 ± 0.334 | 0.437 ± 0.242 |

| Lipid species (nmol/mg DNA) | Lean | OBNG | OBIG |
|--------------------------------|------------------|------------------|-----------------|
| SM - total | 113.433 ± 32.453 | 100.349 ± 38.239 | 76.856 ± 30.385 |
| PE 32:0 | 0.274 ± 0.093 | 0.209 ± 0.065 | 0.206 ± 0.062 |
| PE 34:0 | 0.09 ± 0.058 | 0.068 ± 0.039 | 0.085 ± 0.038 |
| PE 36:0 | 0.224 ± 0.133 | 0.23 ± 0.139 | 0.242 ± 0.135 |
| PE 38:0 | 0.469 ± 0.169 | 0.426 ± 0.165 | 0.44 ± 0.192 |
| PE 40:0 | 0.125 ± 0.046 | 0.105 ± 0.049 | 0.085 ± 0.037 |
| PE 26:1 | 0.574 ± 1.176 | 0.21 ± 0.163 | 0.187 ± 0.304 |
| PE 28:1 | 0.396 ± 0.827 | 0.137 ± 0.111 | 0.111 ± 0.16 |
| PE 32:1 | 0.156 ± 0.059 | 0.145 ± 0.058 | 0.122 ± 0.051 |
| PE 34:1 | 5.754 ± 2.254 | 4.503 ± 1.55 | 3.813 ± 1.386 |
| PE 36:1 | 10.731 ± 2.062 | 8.492 ± 2.544 | 8.215 ± 2.627 |
| PE 38:1 | 1.296 ± 0.755 | 1.128 ± 0.565 | 1.054 ± 0.55 |
| PE 40:1 | 0.301 ± 0.171 | 0.27 ± 0.133 | 0.235 ± 0.131 |
| PE 36:2 | 11.339 ± 2.998 | 8.626 ± 2.492 | 7.45 ± 2.79 |
| PE 38:2 | 1.774 ± 0.68 | 1.423 ± 0.553 | 1.23 ± 0.491 |
| PE 40:2 | 0.353 ± 0.145 | 0.308 ± 0.124 | 0.292 ± 0.139 |
| PE 42:2 | 0.077 ± 0.037 | 0.061 ± 0.022 | 0.058 ± 0.025 |
| PE 34:3 | 0.112 ± 0.035 | 0.1 ± 0.042 | 0.094 ± 0.036 |
| PE 36:3 | 3.291 ± 1.062 | 2.546 ± 0.902 | 2.051 ± 0.891 |
| PE 38:3 | 5.065 ± 0.862 | 4.774 ± 1.501 | 3.825 ± 1.17 |
| PE 40:3 | 0.369 ± 0.102 | 0.291 ± 0.135 | 0.26 ± 0.104 |
| PE 42:3 | 0.054 ± 0.016 | 0.041 ± 0.02 | 0.037 ± 0.019 |
| PE 36:4 | 7.528 ± 2.467 | 5.901 ± 2.573 | 4.748 ± 1.943 |
| PE 38:4 | 53.687 ± 10.847 | 43.909 ± 14.386 | 37.292 ± 12.511 |
| PE 40:4 | 6.675 ± 1.757 | 5.505 ± 1.765 | 4.623 ± 1.761 |
| PE 42:4 | 0.203 ± 0.05 | 0.181 ± 0.072 | 0.131 ± 0.05 |
| PE 36:5 | 0.427 ± 0.124 | 0.348 ± 0.125 | 0.297 ± 0.115 |
| PE 38:5 | 12.397 ± 3.527 | 9.752 ± 3.756 | 7.797 ± 3.088 |
| PE 40:5 | 7.785 ± 1.43 | 6.713 ± 1.889 | 5.862 ± 1.964 |
| PE 42:5 | 0.356 ± 0.096 | 0.299 ± 0.123 | 0.219 ± 0.084 |
| PE 38:6 | 3.29 ± 0.944 | 2.439 ± 0.841 | 2.111 ± 0.809 |
| PE 40:6 | 6.563 ± 1.233 | 5.362 ± 1.581 | 4.892 ± 1.677 |
| PE 42:6 | 0.321 ± 0.093 | 0.272 ± 0.113 | 0.198 ± 0.075 |
| PE 38:7 | 1.303 ± 0.441 | 1.011 ± 0.417 | 0.813 ± 0.327 |
| PE 40:7 | 10.991 ± 2.636 | 9.071 ± 3.276 | 7.494 ± 2.688 |
| PE 42:7 | 0.799 ± 0.224 | 0.671 ± 0.245 | 0.531 ± 0.208 |
| PE 40:8 | 1.866 ± 0.53 | 1.479 ± 0.631 | 1.175 ± 0.477 |
| PE 42:8 | 0.733 ± 0.174 | 0.639 ± 0.24 | 0.54 ± 0.206 |
| PE 42:9 | 0.933 ± 0.243 | 0.758 ± 0.283 | 0.67 ± 0.24 |

| Lipid species (nmol/mg DNA) | Lean | OBNG | OBIG |
|--------------------------------|-------------------------|-------------------------|-------------------------|
| PE 42:10 | 0.32 ± 0.101 | 0.248 ± 0.108 | 0.207 ± 0.086 |
| PE 42:11 | 0.147 ± 0.049 | 0.11 ± 0.052 | 0.092 ± 0.047 |
| PE 44:11 | 0.061 ± 0.019 | 0.05 ± 0.03 | 0.038 ± 0.027 |
| PE 42:12 | 0.204 ± 0.072 | 0.175 ± 0.061 | 0.154 ± 0.072 |
| PE - total | 159.412 ± 32.413 | 128.985 ± 40.019 | 109.981 ± 36.226 |
| PI 34:1 | 0.912 ± 0.583 | 0.945 ± 0.361 | 0.87 ± 0.348 |
| PI 36:1 | 2.199 ± 1.233 | 1.954 ± 0.648 | 1.844 ± 0.739 |
| PI 34:2 | 0.441 ± 0.296 | 0.456 ± 0.19 | 0.356 ± 0.15 |
| PI 36:2 | 3.289 ± 1.24 | 2.99 ± 0.962 | 2.895 ± 1.036 |
| PI 38:2 | 0.232 ± 0.115 | 0.222 ± 0.1 | 0.198 ± 0.119 |
| PI 36:3 | 0.886 ± 0.346 | 0.828 ± 0.329 | 0.674 ± 0.22 |
| PI 38:3 | 4.755 ± 1.298 | 4.342 ± 1.399 | 3.704 ± 1.162 |
| PI 36:4 | 3.028 ± 0.851 | 2.956 ± 1.297 | 2.377 ± 0.754 |
| PI 38:4 | 60.881 ± 11.41 | 49.586 ± 17.554 | 42.608 ± 13.404 |
| PI 40:4 | 1.132 ± 0.207 | 0.855 ± 0.257 | 0.843 ± 0.292 |
| PI 38:5 | 5.652 ± 1.464 | 4.523 ± 2.132 | 3.513 ± 1.292 |
| PI 40:5 | 1.262 ± 0.183 | 0.998 ± 0.438 | 0.911 ± 0.308 |
| PI 38:6 | 0.315 ± 0.153 | 0.252 ± 0.171 | 0.189 ± 0.09 |
| PI 40:6 | 0.58 ± 0.196 | 0.391 ± 0.184 | 0.331 ± 0.112 |
| PI - total | 85.561 ± 15.651 | 71.298 ± 24.366 | 61.313 ± 18.898 |
| PS 32:0 | 0.122 ± 0.036 | 0.148 ± 0.162 | 0.114 ± 0.054 |
| PS 34:0 | 0.46 ± 0.141 | 0.443 ± 0.252 | 0.441 ± 0.197 |
| PS 34:1 | 1.565 ± 0.344 | 1.651 ± 1.086 | 1.309 ± 0.469 |
| PS 36:1 | 49.634 ± 15.37 | 48.357 ± 32.515 | 35.849 ± 14.231 |
| PS 38:1 | 2.065 ± 0.771 | 2.087 ± 1.621 | 1.221 ± 0.593 |
| PS 40:1 | 0.632 ± 0.27 | 0.638 ± 0.624 | 0.322 ± 0.19 |
| PS 34:2 | 0.168 ± 0.04 | 0.161 ± 0.062 | 0.142 ± 0.056 |
| PS 36:2 | 7.008 ± 1.839 | 7.081 ± 4.446 | 5.312 ± 2.04 |
| PS 38:2 | 2.305 ± 0.788 | 2.37 ± 1.726 | 1.494 ± 0.548 |
| PS 40:2 | 0.809 ± 0.331 | 0.809 ± 0.67 | 0.478 ± 0.21 |
| PS 36:3 | 0.552 ± 0.233 | 0.539 ± 0.306 | 0.4 ± 0.155 |
| PS 38:3 | 9.18 ± 3.136 | 10.447 ± 7.02 | 7.138 ± 2.497 |
| PS 40:3 | 0.29 ± 0.107 | 0.271 ± 0.168 | 0.166 ± 0.074 |
| PS 36:4 | 0.365 ± 0.109 | 0.442 ± 0.367 | 0.303 ± 0.12 |
| PS 38:4 | 34.541 ± 11.735 | 34.161 ± 25.209 | 21.523 ± 10.938 |
| PS 40:4 | 5.644 ± 1.645 | 5.713 ± 3.424 | 4.531 ± 1.779 |
| PS 38:5 | 2.119 ± 0.742 | 2.057 ± 1.495 | 1.358 ± 0.656 |
| PS 40:5 | 6.895 ± 1.869 | 7.157 ± 3.511 | 6.094 ± 2.087 |
| PS 42:5 | 0.27 ± 0.106 | 0.263 ± 0.198 | 0.152 ± 0.053 |

| Lipid species (nmol/mg DNA) | Lean | OBNG | OBIG |
|--------------------------------|-------------------------|-------------------------|------------------------|
| PS 38:6 | 0.244 ± 0.056 | 0.278 ± 0.22 | 0.191 ± 0.086 |
| PS 40:6 | 6.329 ± 1.758 | 6.065 ± 3.068 | 5.07 ± 2.019 |
| PS 38:7 | 0.019 ± 0.028 | 0.025 ± 0.04 | 0.015 ± 0.014 |
| PS 40:7 | 0.77 ± 0.345 | 0.849 ± 0.689 | 0.538 ± 0.27 |
| PS 42:7 | 0.118 ± 0.048 | 0.116 ± 0.062 | 0.074 ± 0.029 |
| PS 42:8 | 0.134 ± 0.091 | 0.124 ± 0.089 | 0.094 ± 0.046 |
| PS 42:9 | 0.18 ± 0.143 | 0.136 ± 0.066 | 0.116 ± 0.077 |
| PS - total | 132.415 ± 35.789 | 132.388 ± 86.728 | 94.445 ± 35.832 |
| Cer(d18:1/16:0) | 0.0246 ± 0.0061 | 0.0230 ± 0.0066 | 0.0199 ± 0.0076 |
| Cer(d18:1/18:0) | 0.0039 ± 0.0023 | 0.0034 ± 0.0021 | 0.0024 ± 0.0012 |
| Cer(d18:1/20:0) | 0.0066 ± 0.0035 | 0.0059 ± 0.0041 | 0.0038 ± 0.0022 |
| Cer(d18:1/22:0) | 0.0174 ± 0.0077 | 0.0132 ± 0.0075 | 0.0100 ± 0.0059 |
| Cer(d18:1/24:0) | 0.017 ± 0.0058 | 0.0115 ± 0.0046 | 0.0110 ± 0.0067 |
| Cer(d18:1/24:1(15Z)) | 0.0179 ± 0.005 | 0.0134 ± 0.0044 | 0.0127 ± 0.0041 |
| Cer(d18:1/24:2) | 0.0018 ± 0.0004 | 0.0013 ± 0.0005 | 0.0014 ± 0.0006 |
| Cer(d18:1/24:3) | 0.0013 ± 0.0005 | 0.0009 ± 0.0004 | 0.0009 ± 0.0005 |
| Cer(d18:1/14:4) | 0.0001 ± 0.0001 | 0.0001 ± 0.0001 | 0.0001 ± 0.0001 |
| Cer(d18:1/20:4) | 0.0003 ± 0.0001 | 0.0002 ± 0.0001 | 0.0003 ± 0.0002 |
| Cer(d18:1/24:4) | 0.0004 ± 0.0002 | 0.0003 ± 0.0002 | 0.0003 ± 0.0004 |
| Cer(d18:1/24:5) | 0.0007 ± 0.0003 | 0.0004 ± 0.0003 | 0.0004 ± 0.0011 |
| Cer(d18:1/24:7) | 0.0028 ± 0.0011 | 0.0015 ± 0.0007 | 0.0015 ± 0.0007 |
| Cer(d18:1/16:9) | 0.0019 ± 0.0009 | 0.0016 ± 0.0007 | 0.0020 ± 0.0008 |
| Cer(d18:1/22:9) | 0.0013 ± 0.0007 | 0.0008 ± 0.0004 | 0.0008 ± 0.0008 |
| Cer(d18:1/24:9) | 0.0013 ± 0.0006 | 0.0008 ± 0.0004 | 0.0009 ± 0.0007 |
| Cer - total | 0.099 ± 0.03 | 0.078 ± 0.029 | 0.068 ± 0.027 |
| LPC 14:0 | 0.209 ± 0.103 | 0.159 ± 0.085 | 0.104 ± 0.06 |
| LPC 16:0 | 24.589 ± 11.782 | 18.543 ± 8.17 | 13.798 ± 5.938 |
| LPC 18:0 | 13.248 ± 6.046 | 9.63 ± 3.984 | 7.201 ± 3.152 |
| LPC 20:0 | 0.796 ± 0.322 | 0.62 ± 0.339 | 0.383 ± 0.243 |
| LPC 22:0 | 0.489 ± 0.489 | 0.403 ± 0.315 | 0.255 ± 0.251 |
| LPC 16:1 | 0.758 ± 0.44 | 0.674 ± 0.34 | 0.441 ± 0.204 |
| LPC 18:1 | 23.59 ± 12.172 | 18.39 ± 8.623 | 12.854 ± 5.744 |
| LPC 20:1 | 1.258 ± 0.579 | 0.943 ± 0.427 | 0.627 ± 0.307 |
| LPC 22:1 | 0.474 ± 0.419 | 0.41 ± 0.298 | 0.267 ± 0.24 |
| LPC 16:2 | 0.038 ± 0.02 | 0.039 ± 0.022 | 0.027 ± 0.014 |
| LPC 18:2 | 9.793 ± 5.272 | 6.805 ± 2.88 | 4.928 ± 2.411 |
| LPC 20:2 | 1.019 ± 0.566 | 0.765 ± 0.308 | 0.589 ± 0.267 |
| LPC 22:2 | 0.432 ± 0.486 | 0.379 ± 0.321 | 0.252 ± 0.267 |
| LPC 18:3 | 0.217 ± 0.112 | 0.177 ± 0.093 | 0.109 ± 0.057 |

| Lipid species (nmol/mg DNA) | Lean | OBNG | OBIG |
|--------------------------------|-------------------------|------------------------|------------------------|
| LPC 20:3 | 2.716 ± 1.504 | 2.226 ± 0.925 | 1.509 ± 0.635 |
| LPC 22:3 | 0.536 ± 0.543 | 0.486 ± 0.371 | 0.319 ± 0.291 |
| LPC 20:4 | 17.78 ± 8.975 | 12.787 ± 5.67 | 9.613 ± 4.925 |
| LPC 22:4 | 1.461 ± 0.95 | 1.179 ± 0.638 | 0.808 ± 0.553 |
| LPC 22:5 | 1.781 ± 1.169 | 1.367 ± 0.797 | 1.005 ± 0.685 |
| LPC - total | 101.184 ± 49.085 | 75.983 ± 32.592 | 55.091 ± 24.725 |
| LPE 16:0 | 1.339 ± 0.555 | 0.978 ± 0.539 | 0.627 ± 0.436 |
| LPE 18:0 | 5.685 ± 2.546 | 4.648 ± 2.117 | 3.275 ± 1.708 |
| LPE 16:1 | 0.21 ± 0.225 | 0.157 ± 0.137 | 0.089 ± 0.112 |
| LPE 18:1 | 5.362 ± 3.405 | 4.072 ± 3.056 | 2.411 ± 2.18 |
| LPE 20:1 | 0.313 ± 0.219 | 0.243 ± 0.175 | 0.139 ± 0.129 |
| LPE 18:2 | 1.722 ± 1.114 | 1.223 ± 0.95 | 0.772 ± 0.858 |
| LPE 20:2 | 0.231 ± 0.128 | 0.172 ± 0.101 | 0.099 ± 0.067 |
| LPE 20:3 | 1.73 ± 1.994 | 1.142 ± 0.689 | 0.674 ± 0.865 |
| LPE 22:3 | 2.029 ± 2.845 | 1.062 ± 0.73 | 0.637 ± 0.962 |
| LPE 20:4 | 16.055 ± 13.963 | 10.594 ± 10.371 | 5.474 ± 8.506 |
| LPE 22:4 | 9.928 ± 8.792 | 7.2 ± 7.895 | 3.621 ± 6.351 |
| LPE 20:5 | 0.384 ± 0.315 | 0.314 ± 0.295 | 0.161 ± 0.204 |
| LPE 22:5 | 8.14 ± 6.654 | 5.816 ± 6.357 | 3.136 ± 4.757 |
| LPE - total | 53.129 ± 39.071 | 37.621 ± 31.351 | 21.114 ± 25.871 |
| LPI 16:0 | 0.23 ± 0.281 | 0.137 ± 0.168 | 0.109 ± 0.081 |
| LPI 18:0 | 1.174 ± 0.665 | 0.991 ± 0.583 | 0.667 ± 0.235 |
| LPI 18:1 | 0.156 ± 0.123 | 0.145 ± 0.094 | 0.088 ± 0.05 |
| LPI 20:3 | 0.109 ± 0.07 | 0.089 ± 0.075 | 0.067 ± 0.039 |
| LPI 20:4 | 1.062 ± 0.65 | 0.965 ± 0.657 | 0.592 ± 0.235 |
| LPI - total | 2.732 ± 1.643 | 2.328 ± 1.52 | 1.523 ± 0.54 |
| LPS 16:0 | 0.107 ± 0.095 | 0.261 ± 0.612 | 0.094 ± 0.072 |
| LPS 18:0 | 3.696 ± 2.111 | 3.703 ± 4.166 | 2.055 ± 0.71 |
| LPS 20:0 | 0.092 ± 0.07 | 0.063 ± 0.077 | 0.031 ± 0.015 |
| LPS 18:1 | 2.379 ± 1.382 | 2.446 ± 3.15 | 1.296 ± 0.483 |
| LPS 20:1 | 0.065 ± 0.044 | 0.045 ± 0.049 | 0.024 ± 0.011 |
| LPS 18:2 | 0.137 ± 0.103 | 0.142 ± 0.172 | 0.087 ± 0.038 |
| LPS 20:2 | 0.031 ± 0.018 | 0.028 ± 0.025 | 0.018 ± 0.01 |
| LPS 20:3 | 0.288 ± 0.177 | 0.288 ± 0.322 | 0.17 ± 0.074 |
| LPS 20:4 | 0.9 ± 0.654 | 0.722 ± 0.632 | 0.435 ± 0.206 |
| LPS 22:4 | 0.098 ± 0.043 | 0.081 ± 0.056 | 0.064 ± 0.024 |
| LPS 22:5 | 0.116 ± 0.06 | 0.102 ± 0.075 | 0.077 ± 0.029 |
| LPS - total | 7.909 ± 4.583 | 7.881 ± 9.253 | 4.35 ± 1.502 |

Table S3. Relative abundance of each lipid class. Data are mean \pm SD. Lean vs. OBNG or OBDysG * $p \leq 0.05$. Kruskal-Wallis followed by Dunn post hoc test was performed on data.

| | Overall p-value | Lean | OBNG | OBDysG |
|-------|-----------------|-------------------|-------------------|-------------------|
| % PC | 0.188 | 51.60 \pm 3.87 | 51.06 \pm 3.44 | 52.97 \pm 2.52 |
| % SM | 0.393 | 8.33 \pm 1.38 | 8.89 \pm 1.12 | 8.43 \pm 0.88 |
| % PE | 0.105 | 11.90 \pm 1.68 | 11.59 \pm 1.03 | 12.33 \pm 0.86 |
| % PI | 0.055 | 6.38 \pm 0.66 | 6.38 \pm 0.89 | 6.93 \pm 0.84 |
| % PS | 0.414 | 9.76 \pm 1.85 | 11.31 \pm 4.08 | 10.48 \pm 2.42 |
| % Cer | 0.33 | 0.007 \pm 0.001 | 0.007 \pm 0.001 | 0.008 \pm 0.003 |
| % LPC | 0.245 | 7.46 \pm 3.70 | 6.68 \pm 1.29 | 5.99 \pm 1.25 |
| % LPE | 0.012 | 3.80 \pm 2.22 | 3.24 \pm 2.27 | 2.18 \pm 2.02* |
| % LPI | 0.627 | 0.19 \pm 0.09 | 0.20 \pm 0.08 | 0.18 \pm 0.05 |
| % LPS | 0.704 | 0.56 \pm 0.29 | 0.64 \pm 0.52 | 0.50 \pm 0.14 |

Table S4. Univariable and multivariable associations between lipid classes or species and BMI, with or without adjustment for age.

Data presented as β -coefficients and corresponding 95% confidence intervals (CI). All lipids were log-transformed to the base 10 prior to analyses, and all p-values represent significance of associations after correcting for multiple comparisons using the Benjamini-Hochberg method (bold values indicate $p < 0.05$ after correction). Adjusted β : Multivariable models adjusted for age.

| BMI | | | | | | |
|---------------|-----------------------------|--------------|--------------------------|---------------------------|--------------|--------------------------|
| Lipid class | Unadjusted β (95% CI) | p | R ² (p-value) | Adjusted β (95% CI) | p | R ² (p-value) |
| PC | -0.504 (-0.22, -0.79) | 0.012 | 0.165 (0.001) | -0.418 (-0.13, -0.71) | 0.055 | 0.213 (0.001) |
| | -0.489 (-0.20, -0.77) | 0.012 | 0.157 (0.001) | -0.410 (-0.12, -0.70) | 0.055 | 0.196 (0.001) |
| | -0.386 (-0.09, -0.68) | 0.033 | 0.092 (0.013) | -0.309 (-0.01, -0.61) | 0.119 | 0.127 (0.010) |
| | -0.406 (-0.01, -0.81) | 0.078 | 0.050 (0.052) | -0.364 (0.06, -0.78) | 0.166 | 0.041 (0.120) |
| | -0.419 (-0.06, -0.78) | 0.048 | 0.070 (0.026) | -0.309 (0.05, -0.67) | 0.168 | 0.127 (0.010) |
| | -0.530 (-0.19, -0.87) | 0.018 | 0.131 (0.003) | -0.46 (-0.11, -0.81) | 0.062 | 0.146 (0.005) |
| | -0.587 (-0.14, -1.04) | 0.033 | 0.090 (0.013) | -0.471 (-0.01, -0.93) | 0.119 | 0.125 (0.010) |
| | -1.114 (-0.31, -1.92) | 0.029 | 0.102 (0.009) | -0.823 (-0.02, -1.62) | 0.119 | 0.185 (0.002) |
| | -0.454 (0.01, -0.92) | 0.091 | 0.045 (0.062) | -0.392 (0.09, -0.88) | 0.188 | 0.041 (0.119) |
| | -0.372 (0.16, -0.90) | 0.214 | 0.015 (0.176) | -0.351 (0.21, -0.91) | 0.289 | -0.002 (0.390) |
| Lipid species | Unadjusted β (95% CI) | p | R ² (p-value) | Adjusted β (95% CI) | p | R ² (p-value) |
| PC 28:0 | -0.919 (-0.38, -1.45) | 0.012 | 0.166 (0.001) | -0.858 (-0.29, -1.42) | 0.052 | 0.157 (0.005) |
| SM d18:1/20:1 | -0.624 (-0.21, -1.04) | 0.024 | 0.119 (0.005) | -0.538 (-0.11, -0.97) | 0.079 | 0.134 (0.008) |
| SM d18:1/22:0 | -0.634 (-0.17, -1.10) | 0.029 | 0.099 (0.010) | -0.517 (-0.04, -0.99) | 0.110 | 0.131 (0.008) |
| LPC 20:0 | -1.016 (-0.47, -1.57) | 0.011 | 0.178 (0.001) | -0.908 (-0.34, -1.48) | 0.052 | 0.190 (0.001) |
| LPC 20:3 | -0.559 (-0.10, -1.02) | 0.042 | 0.076 (0.021) | -0.478 (0.00, -0.96) | 0.120 | 0.083 (0.036) |
| LPE 16:0 | -0.849 (-0.29, -1.41) | 0.021 | 0.123 (0.004) | -0.636 (-0.08, -1.19) | 0.096 | 0.214 (0.001) |
| LPE 18:1 | -0.842 (-0.15, -1.54) | 0.042 | 0.077 (0.021) | -0.572 (0.11, -1.25) | 0.170 | 0.179 (0.002) |
| LPE 20:1 | -0.968 (-0.16, -1.78) | 0.044 | 0.074 (0.023) | -0.676 (0.13, -1.48) | 0.170 | 0.159 (0.004) |
| LPE 18:2 | -1.004 (-0.25, -1.76) | 0.032 | 0.093 (0.012) | -0.716 (0.03, -1.46) | 0.130 | 0.188 (0.001) |
| LPE 20:2 | -0.889 (-0.09, -1.69) | 0.054 | 0.063 (0.033) | -0.649 (0.16, -1.45) | 0.188 | 0.119 (0.012) |
| LPE 20:3 | -0.914 (-0.10, -1.73) | 0.054 | 0.063 (0.033) | -0.621 (0.19, -1.43) | 0.208 | 0.148 (0.005) |
| LPE 22:3 | -1.178 (-0.25, -2.10) | 0.036 | 0.088 (0.016) | -0.853 (0.08, -1.78) | 0.141 | 0.160 (0.004) |
| LPE 20:4 | -1.477 (-0.45, -2.51) | 0.026 | 0.109 (0.007) | -1.153 (-0.11, -2.19) | 0.105 | 0.168 (0.003) |
| LPE 22:4 | -1.554 (-0.43, -2.68) | 0.029 | 0.101 (0.009) | -1.180 (-0.05, -2.31) | 0.116 | 0.169 (0.002) |
| LPI 18:0 | -0.482 (-0.03, -0.93) | 0.063 | 0.058 (0.040) | -0.440 (0.03, -0.91) | 0.134 | 0.047 (0.102) |
| LPI 20:4 | -0.411 (0.08, -0.90) | 0.141 | 0.030 (0.105) | -0.326 (0.18, -0.83) | 0.280 | 0.037 (0.136) |

Table S5. Univariable and multivariable associations between lipid classes or species and CRP with or without adjustment for age.

Data presented as β -coefficients and corresponding 95% confidence intervals (CI). All lipids were log-transformed to the base 10 prior to analyses, and all p-values represent significance of associations after correcting for multiple comparisons using the Benjamini-Hochberg method (bold values indicate $p < 0.05$ after correction). Adjusted β : Multivariable models adjusted for age.

| CRP | | | | | | |
|---------------|-----------------------------|--------------|--------------------------|---------------------------|--------------|--------------------------|
| Lipid class | Unadjusted β (95% CI) | p | R ² (p-value) | Adjusted β (95% CI) | p | R ² (p-value) |
| PC | -0.138 (-0.06, -0.22) | 0.012 | 0.160 (0.001) | -0.117 (-0.04, -0.19) | 0.035 | 0.250 (>0.001) |
| | -0.138 (-0.06, -0.22) | 0.012 | 0.164 (0.001) | -0.118 (-0.04, -0.20) | 0.035 | 0.241 (>0.001) |
| | -0.129 (-0.05, -0.21) | 0.015 | 0.138 (0.003) | -0.112 (-0.03, -0.19) | 0.040 | 0.190 (0.002) |
| | -0.168 (-0.06, -0.28) | 0.017 | 0.128 (0.004) | -0.158 (-0.05, -0.27) | 0.040 | 0.123 (0.012) |
| | -0.145 (-0.05, -0.24) | 0.019 | 0.118 (0.006) | -0.121 (-0.02, -0.22) | 0.050 | 0.198 (0.001) |
| | -0.129 (-0.03, -0.23) | 0.027 | 0.096 (0.012) | -0.110 (-0.01, -0.21) | 0.068 | 0.140 (0.007) |
| | -0.190 (-0.06, -0.32) | 0.017 | 0.125 (0.005) | -0.165 (-0.04, -0.29) | 0.044 | 0.172 (0.003) |
| | -0.256 (-0.02, -0.49) | 0.059 | 0.062 (0.037) | -0.191 (0.03, -0.42) | 0.141 | 0.174 (0.003) |
| | -0.147 (-0.01, -0.28) | 0.056 | 0.064 (0.035) | -0.132 (0.00, -0.27) | 0.102 | 0.066 (0.063) |
| | -0.133 (0.02, -0.28) | 0.112 | 0.036 (0.089) | -0.129 (0.03, -0.28) | 0.150 | 0.019 (0.228) |
| Lipid species | Unadjusted β (95% CI) | p | R ² (p-value) | Adjusted β (95% CI) | p | R ² (p-value) |
| PC 28:0 | -0.137 (0.03, -0.30) | 0.130 | 0.033 (0.108) | -0.109 (0.06, -0.27) | 0.241 | 0.059 (0.086) |
| SM d18:1/20:1 | -0.156 (-0.04, -0.28) | 0.029 | 0.092 (0.014) | -0.133 (-0.01, -0.25) | 0.073 | 0.134 (0.009) |
| SM d18:1/22:0 | -0.182 (-0.05, -0.31) | 0.023 | 0.106 (0.009) | -0.153 (-0.02, -0.28) | 0.059 | 0.167 (0.003) |
| LPC 20:0 | -0.246 (-0.09, -0.41) | 0.017 | 0.131 (0.004) | -0.215 (-0.06, -0.37) | 0.043 | 0.175 (0.002) |
| LPC 20:3 | -0.154 (-0.02, -0.29) | 0.047 | 0.071 (0.027) | -0.135 (0.00, -0.27) | 0.096 | 0.087 (0.035) |
| LPE 16:0 | -0.214 (-0.05, -0.38) | 0.028 | 0.094 (0.013) | -0.167 (-0.01, -0.32) | 0.077 | 0.213 (0.001) |
| LPE 18:1 | -0.232 (-0.03, -0.43) | 0.044 | 0.074 (0.025) | -0.174 (0.01, -0.36) | 0.114 | 0.196 (0.001) |
| LPE 20:1 | -0.297 (-0.07, -0.52) | 0.028 | 0.093 (0.014) | -0.233 (-0.02, -0.45) | 0.078 | 0.203 (0.001) |
| LPE 18:2 | -0.273 (-0.06, -0.49) | 0.033 | 0.086 (0.017) | -0.212 (0.00, -0.42) | 0.092 | 0.201 (0.001) |
| LPE 20:2 | -0.302 (-0.08, -0.53) | 0.025 | 0.100 (0.011) | -0.252 (-0.03, -0.47) | 0.067 | 0.165 (0.003) |
| LPE 20:3 | -0.207 (0.03, -0.44) | 0.113 | 0.035 (0.090) | -0.141 (0.08, -0.37) | 0.261 | 0.151 (0.005) |
| LPE 22:3 | -0.214 (0.06, -0.49) | 0.153 | 0.025 (0.132) | -0.137 (0.13, -0.40) | 0.341 | 0.141 (0.008) |
| LPE 20:4 | -0.285 (0.02, -0.59) | 0.094 | 0.042 (0.072) | -0.208 (0.09, -0.50) | 0.210 | 0.133 (0.009) |
| LPE 22:4 | -0.316 (0.01, -0.65) | 0.089 | 0.045 (0.066) | -0.230 (0.09, -0.55) | 0.202 | 0.144 (0.007) |
| LPI 18:0 | -0.155 (-0.03, -0.28) | 0.037 | 0.081 (0.020) | -0.145 (-0.01, -0.27) | 0.073 | 0.075 (0.049) |
| LPI 20:4 | -0.148 (-0.01, -0.29) | 0.063 | 0.059 (0.041) | -0.131 (0.01, -0.27) | 0.114 | 0.067 (0.061) |

Table S6. Univariable and multivariable associations between lipid classes or species and HOMA-IR with or without adjustment for age.

Data presented as β -coefficients and corresponding 95% confidence intervals (CI). All lipids were log-transformed to the base 10 prior to analyses, and all p-values represent significance of associations after correcting for multiple comparisons using the Benjamini-Hochberg method (bold values indicate $p < 0.05$ after correction). Adjusted β : Multivariable models adjusted for age.

| HOMA-IR | | | | | | |
|---------------|-----------------------------|--------------|--------------------------|---------------------------|--------------|--------------------------|
| Lipid class | Unadjusted β (95% CI) | p | R ² (p-value) | Adjusted β (95% CI) | p | R ² (p-value) |
| PC | -0.179 (-0.09, -0.27) | 0.003 | 0.202 (>0.001) | -0.153 (-0.06, -0.24) | 0.010 | 0.298 (>0.001) |
| | -0.180 (-0.09, -0.27) | 0.003 | 0.208 (>0.001) | -0.155 (-0.07, -0.24) | 0.010 | 0.296 (>0.001) |
| | -0.161 (-0.07, -0.26) | 0.007 | 0.161 (0.002) | -0.139 (-0.05, -0.23) | 0.021 | 0.228 (0.001) |
| | -0.168 (-0.04, -0.30) | 0.027 | 0.091 (0.015) | -0.153 (-0.02, -0.29) | 0.058 | 0.093 (0.031) |
| | -0.164 (-0.05, -0.28) | 0.019 | 0.110 (0.008) | -0.134 (-0.02, -0.25) | 0.054 | 0.198 (0.001) |
| | -0.203 (-0.09, -0.31) | 0.004 | 0.189 (0.001) | -0.181 (-0.07, -0.29) | 0.011 | 0.228 (0.001) |
| | -0.180 (-0.03, -0.33) | 0.039 | 0.076 (0.025) | -0.149 (0.00, -0.30) | 0.091 | 0.123 (0.013) |
| | -0.316 (-0.05, -0.59) | 0.040 | 0.074 (0.026) | -0.248 (0.02, -0.51) | 0.099 | 0.158 (0.005) |
| | -0.169 (-0.01, -0.32) | 0.053 | 0.063 (0.038) | -0.148 (0.01, -0.31) | 0.099 | 0.073 (0.054) |
| | -0.169 (0.01, -0.35) | 0.082 | 0.045 (0.066) | -0.162 (0.02, -0.34) | 0.115 | 0.029 (0.176) |
| Lipid species | Unadjusted β (95% CI) | p | R ² (p-value) | Adjusted β (95% CI) | p | R ² (p-value) |
| PC 28:0 | -0.401 (-0.24, -0.56) | 0.001 | 0.316 (>0.001) | -0.381 (-0.21, -0.55) | 0.003 | 0.313 (>0.001) |
| SM d18:1/20:1 | -0.244 (-0.11, -0.38) | 0.005 | 0.18 (0.001) | -0.217 (-0.08, -0.35) | 0.013 | 0.219 (0.001) |
| SM d18:1/22:0 | -0.236 (-0.09, -0.39) | 0.011 | 0.137 (0.003) | -0.204 (-0.05, -0.35) | 0.033 | 0.190 (0.002) |
| LPC 20:0 | -0.346 (-0.17, -0.53) | 0.003 | 0.200 (>0.001) | -0.313 (-0.13, -0.49) | 0.010 | 0.229 (>0.001) |
| LPC 20:3 | -0.198 (-0.04, -0.35) | 0.027 | 0.092 (0.015) | -0.176 (-0.02, -0.33) | 0.060 | 0.106 (0.021) |
| LPE 16:0 | -0.267 (-0.08, -0.46) | 0.018 | 0.113 (0.007) | -0.215 (-0.03, -0.40) | 0.052 | 0.214 (0.001) |
| LPE 18:1 | -0.228 (0.00, -0.46) | 0.077 | 0.048 (0.060) | -0.165 (0.06, -0.39) | 0.187 | 0.152 (0.006) |
| LPE 20:1 | -0.338 (-0.07, -0.60) | 0.029 | 0.089 (0.016) | -0.268 (-0.01, -0.53) | 0.078 | 0.181 (0.002) |
| LPE 18:2 | -0.322 (-0.07, -0.57) | 0.026 | 0.093 (0.014) | -0.258 (-0.02, -0.50) | 0.073 | 0.180 (0.002) |
| LPE 20:2 | -0.317 (-0.05, -0.58) | 0.038 | 0.078 (0.023) | -0.259 (0.00, -0.52) | 0.090 | 0.137 (0.009) |
| LPE 20:3 | -0.319 (-0.05, -0.59) | 0.039 | 0.075 (0.025) | -0.248 (0.02, -0.51) | 0.099 | 0.170 (0.003) |
| LPE 22:3 | -0.393 (-0.09, -0.70) | 0.027 | 0.096 (0.015) | -0.312 (-0.02, -0.61) | 0.075 | 0.183 (0.003) |
| LPE 20:4 | -0.419 (-0.07, -0.76) | 0.035 | 0.081 (0.021) | -0.344 (0.00, -0.68) | 0.086 | 0.138 (0.008) |
| LPE 22:4 | -0.389 (-0.01, -0.77) | 0.065 | 0.054 (0.049) | -0.301 (0.07, -0.67) | 0.149 | 0.124 (0.013) |
| LPI 18:0 | -0.178 (-0.03, -0.33) | 0.038 | 0.078 (0.023) | -0.163 (-0.01, -0.32) | 0.073 | 0.077 (0.049) |
| LPI 20:4 | -0.139 (0.03, -0.30) | 0.117 | 0.032 (0.103) | -0.114 (0.05, -0.28) | 0.209 | 0.052 (0.097) |

Table S7. Univariable and multivariable associations between lipid classes or species and fasting insulin with or without adjustment for age.

Data presented as β -coefficients and corresponding 95% confidence intervals (CI). All lipids were log-transformed to the base 10 prior to analyses, and all p-values represent significance of associations after correcting for multiple comparisons using the Benjamini-Hochberg method (bold values indicate $p < 0.05$ after correction). Adjusted β : Multivariable models adjusted for age.

| Fasting insulin | | | | | | |
|-----------------|-----------------------------|--------------|--------------------------|---------------------------|--------------|--------------------------|
| Lipid class | Unadjusted β (95% CI) | p | R ² (p-value) | Adjusted β (95% CI) | p | R ² (p-value) |
| PC | -0.198 (-0.09, -0.30) | 0.005 | 0.189 (0.001) | -0.175 (-0.08, -0.27) | 0.009 | 0.303 (>0.001) |
| | -0.198 (-0.09, -0.30) | 0.005 | 0.191 (0.001) | -0.176 (-0.08, -0.27) | 0.009 | 0.298 (>0.001) |
| | -0.180 (-0.07, -0.29) | 0.009 | 0.154 (0.002) | -0.160 (-0.06, -0.26) | 0.018 | 0.235 (>0.001) |
| | -0.186 (-0.04, -0.34) | 0.035 | 0.085 (0.019) | -0.173 (-0.02, -0.32) | 0.058 | 0.093 (0.031) |
| | -0.177 (-0.04, -0.31) | 0.027 | 0.097 (0.013) | -0.151 (-0.02, -0.28) | 0.052 | 0.198 (0.001) |
| | -0.214 (-0.09, -0.34) | 0.008 | 0.159 (0.002) | -0.195 (-0.07, -0.32) | 0.016 | 0.213 (0.001) |
| | -0.195 (-0.02, -0.37) | 0.052 | 0.067 (0.033) | -0.169 (0.00, -0.34) | 0.091 | 0.124 (0.013) |
| | -0.341 (-0.03, -0.65) | 0.055 | 0.065 (0.035) | -0.283 (0.01, -0.58) | 0.100 | 0.160 (0.004) |
| | -0.183 (-0.01, -0.36) | 0.068 | 0.055 (0.048) | -0.165 (0.01, -0.34) | 0.111 | 0.071 (0.057) |
| | -0.181 (0.02, -0.38) | 0.105 | 0.038 (0.085) | -0.173 (0.03, -0.38) | 0.133 | 0.023 (0.205) |
| Lipid species | Unadjusted β (95% CI) | p | R ² (p-value) | Adjusted β (95% CI) | p | R ² (p-value) |
| PC 28:0 | -0.426 (-0.23, -0.62) | 0.003 | 0.262 (>0.001) | -0.401 (-0.20, -0.60) | 0.006 | 0.269 (>0.001) |
| SM d18:1/20:1 | -0.264 (-0.11, -0.42) | 0.008 | 0.161 (0.002) | -0.240 (-0.09, -0.39) | 0.016 | 0.212 (0.001) |
| SM d18:1/22:0 | -0.247 (-0.07, -0.42) | 0.022 | 0.112 (0.008) | -0.218 (-0.05, -0.39) | 0.040 | 0.179 (0.002) |
| LPC 20:0 | -0.366 (-0.16, -0.57) | 0.007 | 0.170 (0.001) | -0.336 (-0.13, -0.54) | 0.013 | 0.213 (0.001) |
| LPC 20:3 | -0.220 (-0.04, -0.40) | 0.034 | 0.086 (0.018) | -0.200 (-0.02, -0.38) | 0.058 | 0.107 (0.021) |
| LPE 16:0 | -0.291 (-0.08, -0.51) | 0.024 | 0.102 (0.011) | -0.247 (-0.04, -0.45) | 0.047 | 0.217 (0.001) |
| LPE 18:1 | -0.241 (0.03, -0.51) | 0.103 | 0.039 (0.083) | -0.188 (0.07, -0.44) | 0.181 | 0.153 (0.005) |
| LPE 20:1 | -0.365 (-0.06, -0.67) | 0.04 | 0.079 (0.023) | -0.306 (-0.02, -0.60) | 0.075 | 0.182 (0.002) |
| LPE 18:2 | -0.354 (-0.07, -0.64) | 0.035 | 0.085 (0.019) | -0.299 (-0.03, -0.57) | 0.066 | 0.184 (0.002) |
| LPE 20:2 | -0.314 (-0.01, -0.62) | 0.068 | 0.055 (0.049) | -0.264 (0.03, -0.56) | 0.118 | 0.125 (0.012) |
| LPE 20:3 | -0.327 (-0.02, -0.64) | 0.065 | 0.057 (0.045) | -0.266 (0.03, -0.56) | 0.118 | 0.165 (0.004) |
| LPE 22:3 | -0.428 (-0.08, -0.78) | 0.037 | 0.085 (0.020) | -0.356 (-0.02, -0.69) | 0.073 | 0.185 (0.003) |
| LPE 20:4 | -0.453 (-0.06, -0.85) | 0.048 | 0.071 (0.029) | -0.389 (0.00, -0.77) | 0.086 | 0.139 (0.008) |
| LPE 22:4 | -0.420 (0.01, -0.85) | 0.085 | 0.047 (0.063) | -0.345 (0.07, -0.77) | 0.142 | 0.126 (0.012) |
| LPI 18:0 | -0.194 (-0.02, -0.37) | 0.049 | 0.070 (0.030) | -0.180 (-0.01, -0.35) | 0.077 | 0.074 (0.053) |
| LPI 20:4 | -0.154 (0.03, -0.34) | 0.131 | 0.029 (0.113) | -0.133 (0.05, -0.32) | 0.197 | 0.054 (0.091) |

Table S8. Univariable and multivariable associations between lipid species or classes and Fasting glucose with or without adjustment for age.

Data presented as β -coefficients and corresponding 95% confidence intervals (CI). All lipids were log-transformed to the base 10 prior to analyses, and all p-values represent significance of associations after correcting for multiple comparisons using the Benjamini-Hochberg method (bold values indicate $p < 0.05$ after correction). Adjusted β : Multivariable models adjusted for age.

| Fasting glucose | | | | | | |
|-----------------|-----------------------------|--------------|--------------------------|---------------------------|--------------|--------------------------|
| Lipid class | Unadjusted β (95% CI) | p | R ² (p-value) | Adjusted β (95% CI) | p | R ² (p-value) |
| PC | -0.518 (-0.09, -0.95) | 0.090 | 0.076 (0.021) | -0.350 (0.09, -0.79) | 0.337 | 0.133 (0.008) |
| | -0.527 (-0.10, -0.95) | 0.090 | 0.081 (0.018) | -0.375 (0.07, -0.82) | 0.316 | 0.125 (0.010) |
| | -0.417 (0.01, -0.85) | 0.120 | 0.044 (0.063) | -0.270 (0.18, -0.72) | 0.414 | 0.084 (0.035) |
| | -0.465 (0.12, -1.05) | 0.179 | 0.025 (0.123) | -0.385 (0.24, -1.01) | 0.405 | 0.017 (0.237) |
| | -0.557 (-0.04, -1.08) | 0.095 | 0.058 (0.040) | -0.361 (0.18, -0.90) | 0.391 | 0.110 (0.016) |
| | -0.750 (-0.26, -1.24) | 0.061 | 0.127 (0.004) | -0.639 (-0.12, -1.15) | 0.288 | 0.135 (0.007) |
| | -0.600 (0.06, -1.26) | 0.138 | 0.037 (0.081) | -0.372 (0.32, -1.06) | 0.465 | 0.079 (0.041) |
| | -1.155 (0.04, -2.35) | 0.120 | 0.045 (0.062) | -0.593 (0.61, -1.80) | 0.488 | 0.138 (0.007) |
| | -0.492 (0.18, -1.17) | 0.209 | 0.018 (0.159) | -0.373 (0.35, -1.10) | 0.465 | 0.016 (0.244) |
| | -0.528 (0.23, -1.29) | 0.221 | 0.015 (0.180) | -0.502 (0.32, -1.32) | 0.408 | -0.003 (0.404) |
| Lipid species | Unadjusted β (95% CI) | p | R ² (p-value) | Adjusted β (95% CI) | p | R ² (p-value) |
| PC 28:0 | -1.586 (-0.87, -2.30) | 0.015 | 0.258 (>0.001) | -1.557 (-0.78, -2.33) | 0.054 | 0.243 (>0.001) |
| SM d18:1/20:1 | -0.809 (-0.20, -1.42) | 0.084 | 0.094 (0.012) | -0.661 (-0.02, -1.3) | 0.304 | 0.106 (0.018) |
| SM d18:1/22:0 | -0.941 (-0.28, -1.60) | 0.079 | 0.108 (0.007) | -0.750 (-0.05, -1.45) | 0.304 | 0.130 (0.009) |
| LPC 20:0 | -1.263 (-0.45, -2.07) | 0.061 | 0.130 (0.003) | -1.073 (-0.22, -1.93) | 0.288 | 0.139 (0.007) |
| LPC 20:3 | -0.582 (0.09, -1.26) | 0.158 | 0.032 (0.097) | -0.423 (0.29, -1.14) | 0.422 | 0.042 (0.118) |
| LPE 16:0 | -0.897 (-0.07, -1.73) | 0.095 | 0.059 (0.039) | -0.489 (0.35, -1.32) | 0.426 | 0.160 (0.003) |
| LPE 18:1 | -0.932 (0.08, -1.95) | 0.132 | 0.039 (0.077) | -0.421 (0.60, -1.44) | 0.533 | 0.148 (0.005) |
| LPE 20:1 | -1.170 (0.01, -2.35) | 0.116 | 0.048 (0.056) | -0.632 (0.56, -1.83) | 0.465 | 0.134 (0.008) |
| LPE 18:2 | -1.076 (0.04, -2.19) | 0.120 | 0.044 (0.064) | -0.525 (0.60, -1.65) | 0.504 | 0.148 (0.005) |
| LPE 20:2 | -1.476 (-0.35, -2.60) | 0.085 | 0.092 (0.013) | -1.088 (0.08, -2.26) | 0.309 | 0.131 (0.008) |
| LPE 20:3 | -1.423 (-0.26, -2.59) | 0.090 | 0.078 (0.020) | -0.925 (0.27, -2.12) | 0.340 | 0.150 (0.005) |
| LPE 22:3 | -1.435 (-0.09, -2.78) | 0.096 | 0.059 (0.042) | -0.874 (0.50, -2.25) | 0.396 | 0.134 (0.009) |
| LPE 20:4 | -1.513 (0.02, -3.04) | 0.116 | 0.047 (0.058) | -0.879 (0.69, -2.45) | 0.448 | 0.115 (0.014) |
| LPE 22:4 | -1.422 (0.25, -3.10) | 0.162 | 0.031 (0.102) | -0.671 (1.03, -2.38) | 0.538 | 0.114 (0.014) |
| LPI 18:0 | -0.506 (0.15, -1.16) | 0.190 | 0.023 (0.136) | -0.421 (0.28, -1.12) | 0.414 | 0.013 (0.263) |
| LPI 20:4 | -0.341 (0.37, -1.05) | 0.389 | -0.002 (0.352) | -0.163 (0.59, -0.92) | 0.732 | 0.012 (0.272) |

Table S9. Univariable and multivariable associations between lipid classes or species and HbA1c, with or without adjustment for age.

Data presented as β -coefficients and corresponding 95% confidence intervals (CI). All lipids were log-transformed to the base 10 prior to analyses, and all p-values represent significance of associations after correcting for multiple comparisons using the Benjamini-Hochberg method (bold values indicate $p < 0.05$ after correction). Adjusted β : Multivariable models adjusted for age.

| HbA1c | | | | | | |
|---------------|-----------------------------|--------------|--------------------------|---------------------------|-------|--------------------------|
| Lipid class | Unadjusted β (95% CI) | p | R ² (p-value) | Adjusted β (95% CI) | p | R ² (p-value) |
| PC | -0.968 (-0.28, -1.66) | 0.044 | 0.105 (0.008) | -0.731 (-0.03, -1.43) | 0.180 | 0.159 (0.003) |
| | -0.967 (-0.28, -1.65) | 0.044 | 0.107 (0.008) | -0.750 (-0.05, -1.45) | 0.180 | 0.150 (0.005) |
| | -0.706 (0.00, -1.41) | 0.093 | 0.049 (0.054) | -0.491 (0.23, -1.21) | 0.306 | 0.091 (0.029) |
| | -0.799 (0.15, -1.75) | 0.143 | 0.030 (0.104) | -0.680 (0.32, -1.68) | 0.306 | 0.022 (0.205) |
| | -1.103 (-0.27, -1.93) | 0.051 | 0.094 (0.012) | -0.833 (0.02, -1.68) | 0.186 | 0.140 (0.006) |
| | -1.250 (-0.46, -2.04) | 0.037 | 0.133 (0.003) | -1.079 (-0.25, -1.91) | 0.144 | 0.144 (0.006) |
| | -1.062 (0.01, -2.14) | 0.094 | 0.047 (0.058) | -0.733 (0.38, -1.84) | 0.317 | 0.088 (0.031) |
| | -2.134 (-0.21, -4.06) | 0.071 | 0.062 (0.034) | -1.333 (0.59, -3.26) | 0.305 | 0.152 (0.004) |
| | -0.663 (0.45, -1.77) | 0.291 | 0.006 (0.248) | -0.467 (0.70, -1.64) | 0.508 | 0.008 (0.299) |
| | -0.746 (0.50, -2.00) | 0.291 | 0.006 (0.248) | -0.686 (0.64, -2.01) | 0.402 | -0.010 (0.495) |
| Lipid species | Unadjusted β (95% CI) | p | R ² (p-value) | Adjusted β (95% CI) | p | R ² (p-value) |
| PC 28:0 | -2.444 (-1.28, -3.61) | 0.035 | 0.233 (>0.001) | -2.342 (-1.10, -3.58) | 0.123 | 0.222 (0.001) |
| SM d18:1/20:1 | -1.57 (-0.60, -2.54) | 0.037 | 0.140 (0.002) | -1.368 (-0.36, -2.38) | 0.144 | 0.149 (0.005) |
| SM d18:1/22:0 | -1.728 (-0.67, -2.79) | 0.037 | 0.141 (0.002) | -1.458 (-0.36, -2.56) | 0.144 | 0.162 (0.003) |
| LPC 20:0 | -2.221 (-0.92, -3.52) | 0.037 | 0.154 (0.002) | -1.942 (-0.58, -3.30) | 0.144 | 0.164 (0.003) |
| LPC 20:3 | -1.081 (0.02, -2.18) | 0.094 | 0.047 (0.059) | -0.856 (0.29, -2.00) | 0.272 | 0.055 (0.080) |
| LPE 16:0 | -1.649 (-0.31, -2.99) | 0.056 | 0.079 (0.019) | -1.067 (0.27, -2.40) | 0.239 | 0.177 (0.002) |
| LPE 18:1 | -1.686 (-0.04, -3.33) | 0.088 | 0.052 (0.049) | -0.954 (0.68, -2.58) | 0.357 | 0.158 (0.004) |
| LPE 20:1 | -1.864 (0.06, -3.79) | 0.098 | 0.044 (0.063) | -1.065 (0.86, -2.99) | 0.374 | 0.135 (0.007) |
| LPE 18:2 | -1.996 (-0.19, -3.80) | 0.071 | 0.062 (0.034) | -1.212 (0.58, -3.00) | 0.306 | 0.162 (0.003) |
| LPE 20:2 | -2.389 (-0.55, -4.22) | 0.051 | 0.090 (0.014) | -1.802 (0.08, -3.69) | 0.191 | 0.134 (0.008) |
| LPE 20:3 | -2.423 (-0.53, -4.31) | 0.052 | 0.087 (0.015) | -1.692 (0.22, -3.60) | 0.210 | 0.160 (0.003) |
| LPE 22:3 | -2.333 (-0.13, -4.54) | 0.080 | 0.058 (0.043) | -1.498 (0.72, -3.71) | 0.306 | 0.137 (0.008) |
| LPE 20:4 | -2.676 (-0.20, -5.16) | 0.077 | 0.058 (0.039) | -1.762 (0.75, -4.28) | 0.300 | 0.126 (0.010) |
| LPE 22:4 | -2.587 (0.13, -5.30) | 0.103 | 0.042 (0.067) | -1.513 (1.22, -4.25) | 0.374 | 0.124 (0.011) |
| LPI 18:0 | -0.710 (0.36, -1.79) | 0.246 | 0.012 (0.200) | -0.567 (0.57, -1.70) | 0.419 | 0.005 (0.325) |
| LPI 20:4 | -0.469 (0.70, -1.63) | 0.470 | -0.007 (0.434) | -0.198 (1.02, -1.41) | 0.777 | 0.010 (0.283) |