

SUPPLEMENTAL MATERIAL

Online Supporting Material

Supplemental Figures and Tables:

Table s1- Particle Diameter ranges.

Table s2- Baseline characteristics of 26034 apparently healthy women according to quintiles (Q) of fish and energy adjusted marine n-3 intake.

Table s3- Baseline characteristics of 26034 apparently healthy women according to quintiles (Q) of energy adjusted ALA, EPA and DHA fatty acid intake.

Table s4- Raw means (or geometrical means) and adjusted means of LDL related variables according to quintiles of energy adjusted EPA intake.

Table s5- Raw means (or geometrical means) and adjusted means of LDL related variables according to quintiles of energy adjusted DHA intake.

Table s6- Raw means (or geometrical means) and adjusted means of LDL related variables according to quintiles of energy adjusted ALA intake.

Table s7- Raw means (or geometrical means) and adjusted means of VLDL related variables according to quintiles of energy adjusted EPA intake.

Table s8- Raw means (or geometrical means) and adjusted means of VLDL related variables according to quintiles of energy adjusted DHA intake.

Table s9- Raw means (or geometrical means) and adjusted means of VLDL related variables according to quintiles of energy adjusted ALA intake.

Table s10- Raw means (or geometrical means) and adjusted means of HDL related variables according to quintiles of energy adjusted EPA intake.

Table s11- Raw means (or geometrical means) and adjusted means of HDL related variables according to quintiles of energy adjusted DHA intake.

Table s12- Raw means (or geometrical means) and adjusted means of HDL related variables according to quintiles of energy adjusted ALA intake.

Table s13- Correspondence between absolute values and per 1% and per 5% differences.

Table s14- Adjusted means for the lower exposure quintiles (Q1) that showed a significant association ($P_{trend} < 0.05$).

Table s1- Particle diameter ranges.

| | Diameter range (nm) | % CV |
|----------------|---------------------|------|
| LDL Particles | | |
| Total | 18-23 | 2.1 |
| Large | 21.2-23 | 6.3 |
| Small | 18-21.2 | 4.7 |
| IDL Particles | 23-27 | 13.1 |
| HDL Particles | | |
| Total | 7.3-13 | 1.5 |
| Large | 8.8-13 | 5.9 |
| Medium | 8.2-8.8 | <30 |
| Small | 7.3-8.2 | 3.7 |
| VLDL Particles | | |
| Total | ≥ 27 | 3.1 |
| Large | > 60 | 5.1 |
| Medium | 35-60 | 4.1 |
| Small | 27-35 | 7.1 |

Table s2- Baseline characteristics of 26,034 apparently healthy women according to quintiles of fish and energy adjusted marine n-3 intake.

| Characteristic | Total fish intake (median [min, max] serv/day) | | | | Total marine n-3 (median [min, max] g/day) | | | |
|---|--|--------------------|--------------------|----------|--|--------------------|--------------------|----------|
| | Q:1 | Q:3 | Q:5 | <i>P</i> | Q:1 | Q:3 | Q:5 | <i>P</i> |
| | 0.07 [0, 0.07] | 0.21 [0.2, 0.21] | 0.5 [0.43, 0.64] | | 0.95 [0.86, 1.02] | 1.35 [1.31, 1.39] | 1.89 [1.77, 2.1] | |
| <i>N</i> | 5839 | 5465 | 5617 | | 5248 | 4991 | 5123 | |
| Age (y) | 52.5 [48.6, 59.0] | 52.8 [49.0, 58.8] | 53.2 [49.2, 59.0] | 0.0025 | 52.1 [48.5, 57.7] | 52.7 [48.9, 58.7] | 54 [49.6, 60] | <0.0001 |
| White (%) | 96.0 | 95.8 | 94.0 | <0.0001 | 95.2 | 96.0 | 95.1 | <0.0001 |
| BMI (kg/m ²) | 24.7 [22.3, 28.2] | 24.6 [22.3, 27.8] | 25.0 [22.6, 28.3] | <0.0001 | 24.3 [22.1, 27.9] | 24.9 [22.5, 28.2] | 25.0 [22.6, 28.3] | <0.0001 |
| Exercise (MET-h/wk) | 6.87 [1.97, 17.51] | 9.11 [3.03, 20.74] | 11.3 [4.02, 23.55] | <0.0001 | 7.70 [2.30, 19.5] | 9.20 [3.10, 20.94] | 9.90 [3.01, 21.50] | <0.0001 |
| Fruits and vegetables (serving/d) | 4.44 [2.96, 6.30] | 5.45 [3.93, 7.34] | 6.96 [5.08, 9.32] | <0.0001 | 4.66 [3.14, 6.64] | 5.58 [4.01, 7.63] | 6.06 [4.29, 8.32] | <0.0001 |
| Nuts (serving/d) | 0 [0, 0.13] | 0.07 [0, 0.13] | 0.07 [0, 0.13] | <0.0001 | 0 [0, 0.13] | 0.07 [0, 0.13] | 0.07 [0, 0.13] | <0.0001 |
| Red meat (serving/d) | 0.55 [0.27, 0.91] | 0.63 [0.34, 0.98] | 0.56 [0.33, 0.98] | <0.0001 | 0.63 [0.34, 1.06] | 0.63 [0.34, 0.99] | 0.50 [0.28, 0.85] | <0.0001 |
| Energy intake (kcal/d) | 1471 [1177, 1826] | 1686 [1377, 2035] | 1896 [1557, 2284] | <0.0001 | 1639 [1324, 2008] | 1707 [1368, 2105] | 1649 [1322, 2017] | <0.0001 |
| Dietary magnesium (mg) ¹ | 314 [273, 366] | 327 [289, 372] | 352 [314, 399] | <0.0001 | 323 [279, 374] | 329 [291, 377] | 334 [294, 386] | <0.0001 |
| <i>trans</i> Fatty acids (g/d) ¹ | 2.27 [1.65, 3.09] | 2.10 [1.58, 2.82] | 1.81 [1.33, 2.47] | <0.0001 | 2.02 [1.48, 2.7] | 2.1 [1.56, 2.83] | 2.08 [1.47, 2.86] | <0.0001 |
| Saturated fatty acids (g/d) ¹ | 20.3 [17.1, 23.7] | 19.5 [16.7, 22.5] | 18.0 [15.2, 20.9] | <0.0001 | 19.6 [16.3, 23.1] | 19.4 [16.5, 22.4] | 19.0 [16.3, 21.9] | <0.0001 |
| Cereal fiber (g/d) ¹ | 4.27 [3.57, 5.12] | 4.33 [3.70, 5.11] | 4.57 [3.91, 5.41] | <0.0001 | 4.11 [3.44, 4.91] | 4.4 [3.74, 5.16] | 4.62 [3.92, 5.45] | <0.0001 |
| Parental history of diabetes (%) | 24.5 | 23.9 | 26.3 | 0.01 | 24.3 | 24.6 | 26.0 | 0.13 |
| Glycated hemoglobin (%) | 4.99 [4.84, 5.18] | 4.99 [4.83, 5.17] | 5.00 [4.83, 5.18] | 0.13 | 4.99 [4.83, 5.17] | 4.99 [4.83, 5.18] | 5.00 [4.83, 5.17] | 0.33 |
| Current smoking (%) | 12.5 | 11.6 | 10.2 | <0.0001 | 14.2 | 10.4 | 11.5 | <0.0001 |
| Daily alcohol use (%) | 7.1 | 11.9 | 12.0 | <0.0001 | 11.8 | 10.5 | 9.78 | <0.0001 |
| Postmenopausal (%) | 53.6 | 53.8 | 54.7 | 0.0019 | 51.5 | 53.7 | 57.5 | <0.0001 |
| Hormone Therapy use (%) | 43.0 | 44.5 | 44.5 | 0.03 | 42.2 | 44.3 | 45.8 | <0.001 |
| Hypertension (%) | 23.2 | 23.6 | 25.8 | 0.01 | 23.2 | 23.1 | 26.0 | 0.0015 |

Values shown are medians (25th 75th percentile) or percentages. MET-h indicates metabolic equivalent task hours. P values were derived from Wilcoxon rank sum test (continuous variables) or chi-squared test (categorical variables).¹Energy adjusted.

Table s3- Baseline characteristics of 26034 women according to quintiles (Q) of energy adjusted EPA, DHA and ALA fatty acid intake.

| Characteristic | Eicosapentaenoic acid (EPA) | | | | Docosahexaenoic acid (DHA) | | | | α -Linolenic acid (ALA) | | | |
|---|-----------------------------|----------------------|----------------------|----------|----------------------------|----------------------|---------------------|----------|--------------------------------|----------------------|----------------------|----------|
| | Q:1 | Q:3 | Q:5 | <i>P</i> | Q:1 | Q:3 | Q:5 | <i>P</i> | Q:1 | Q:3 | Q:5 | <i>P</i> |
| <i>N</i> | 5370 | 3226 | 4947 | | 6351 | 4860 | 4797 | | 5286 | 5225 | 5079 | |
| Age (y) | 52.6 [48.6 59.0] | 52.7 [48.8 58.3] | 53.6 [49.4 59.2] | <.0001 | 52.5 [48.6 58.9] | 52.8 [48.8 58.6] | 53.4 [49.5 59.1] | <.0001 | 52.2 [48.6 58.0] | 52.7 [48.9 58.7] | 53.8 [49.4 60.1] | <.0001 |
| White (%) | 96.7 | 96.5 | 92.1 | <.0001 | 96.7 | 96.1 | 92.2 | <.0001 | 93.7 | 96.0 | 96.8 | <.0001 |
| BMI (kg/m²) | 24.9 [22.3 28.3] | 25.1 [22.6 28.3] | 24.5 [22.3 27.5] | <.0001 | 24.8 [22.3 28.3] | 24.8 [22.5 28.3] | 24.8 [22.5 28.2] | 0.57 | 24.3 [22.1 27.8] | 24.9 [22.6 28.3] | 25.0 [22.5 28.4] | <.0001 |
| Exercise (MET-h/wk) | 7.14 [2.11 17.50] | 8.83 [2.92 20.20] | 11.8 [4.25 24.39] | <.0001 | 7.00 [2.12 17.87] | 8.70 [2.87 20.20] | 11.8 [4.15 24.4] | <.0001 | 8.69 [2.63 20.94] | 9.00 [3.01 20.61] | 8.68 [2.64 20.20] | 0.01 |
| Fruits & veg. (serving/d) | 4.92 [3.44 6.82] | 5.36 [3.77 7.30] | 6.04 [4.28 8.23] | <.0001 | 4.89[3.33 6.87] | 5.45 [3.89 7.50] | 6.15 [4.34 8.43] | <.0001 | 4.87 [3.25 6.86] | 5.65 [4.00 7.73] | 5.81 [4.10 7.98] | <.0001 |
| Nuts (serving/d) | 0 [0 0.13] | 0 [0 0.13] | 0.07 [0 0.13] | <.0001 | 0 [0 0.13] | 0.07 [0 0.13] | 0.07 [0 0.13] | <.0001 | 0 [0 0.13] | 0.07 [0 0.13] | 0.07 [0 0.13] | <.0001 |
| Red meat (serving/d) | 0.64 [0.34 1.06] | 0.56 [0.34 0.91] | 0.43 [0.27 0.77] | <.0001 | 0.70 [0.34 1.07] | 0.63 [0.35 0.99] | 0.42 [0.27 0.71] | <.0001 | 0.56 [0.29 0.98] | 0.63 [0.34 0.99] | 0.56 [0.33 0.91] | <.0001 |
| Energy intake (kcal/d) | 1604 [1325 1988] | 1545 [1281 1954] | 1577 [1288 1947] | <.0001 | 1655 [1330 2050] | 1681 [1362 2074] | 1592 [1280 1961] | <.0001 | 1644 [1329 2004] | 1713 [1368 2093] | 1656 [1334 2026] | <.0001 |
| Diet. magnesium (mg)¹ | 314 [274 364] | 331 [294 378] | 351 [312 400] | <.0001 | 312 [272 362] | 327 [291 373] | 357 [316 406] | <.0001 | 332 [285 384] | 331 [291 379] | 323 [286 370] | <.0001 |
| trans FA (g/d)¹ | 2.31 [1.67 3.12] | 2.08 [1.56 2.76] | 1.75 [1.29 2.35] | <.0001 | 2.34 [1.71 3.15] | 2.10 [1.59 2.82] | 1.72 [1.27 2.33] | <.0001 | 1.86 [1.37 2.55] | 2.11 [1.58 2.79] | 2.28 [1.62 3.08] | <.0001 |
| Saturated FA (g/d)¹ | 20.4 [17.2 23.7] | 19.3 [16.5 22.1] | 17.9 [15.1 20.9] | <.0001 | 20.5 [17.3 23.8] | 19.5 [16.7 22.3] | 17.7 [14.9 20.7] | <.0001 | 18.9 [15.6 22.3] | 19.3 [16.6 22.3] | 19.6 [16.9 22.5] | <.0001 |
| Cereal fibre (g/d)¹ | 4.27 [3.59 5.13] | 4.38 [3.75 5.12] | 4.58 [3.89 5.44] | <.0001 | 4.24 [3.57 5.09] | 4.33 [3.70 5.10] | 4.61 [3.92 5.49] | <.0001 | 4.15 [3.47 4.99] | 4.39 [3.73 5.16] | 4.55 [3.86 5.38] | <.0001 |
| Par. Hist. diabetes (%) | 24.4 | 26.0 | 24.8 | 0.17 | 24.4 | 24.2 | 25.8 | 0.32 | 24.4 | 24.7 | 25.6 | 0.38 |

Online Supporting Material

| | | | | | | | | | | | | |
|------------------------------|---------------------|---------------------|---------------------|--------|---------------------|---------------------|---------------------|--------|---------------------|---------------------|---------------------|--------|
| GH (%) | 5.00 [4.84 5.18] | 4.99 [4.83 5.16] | 4.98 [4.82 5.17] | 0.03 | 5.00 [4.84 5.17] | 4.99 [4.84 5.17] | 4.99 [4.83 5.17] | 0.25 | 4.99 [4.83 5.17] | 4.98 [4.83 5.17] | 5.01 [4.84 5.18] | 0.03 |
| Current smoking (%) | 12.4 | 13.2 | 10.1 | <.0001 | 12.9 | 11.9 | 10.3 | <.0001 | 13.8 | 10.4 | 12.1 | <.0001 |
| Daily alcohol use (%) | 7.32 | 10.3 | 13.7 | <.0001 | 8.36 | 10.3 | 11.5 | <.0001 | 13.5 | 9.90 | 9.18 | <.0001 |
| Postmenopausal (%) | 53.7 | 52.6 | 55.8 | <.0001 | 53.5 | 54.3 | 55.4 | <.0001 | 51.6 | 54.1 | 57.6 | <.0001 |
| HT use (%) | 42.4 | 45.2 | 46.8 | <.0001 | 42.4 | 43.6 | 46.0 | <.001 | 41.7 | 44.4 | 45.5 | 0.0018 |
| Hypertension (%) | 22.9 | 24.0 | 24.4 | 0.22 | 23.3 | 23.7 | 25.8 | 0.02 | 23.6 | 23.6 | 25.4 | 0.14 |

Values shown are medians (25th 75th percentile) or percentages. MET-h indicates metabolic equivalent task hours. veg.: vegetables. Diet. Magnesium: Dietary magnesium. FA: Fatty Acids. Par. Hist. Diabetes: Parental History of Diabetes. GH: Glycated Haemoglobin. P values were derived from Wilcoxon rank sum test (continuous variables) or chi-squared test (categorical variables). ¹Energy adjusted

Table s4- LDL related variables. Strategy based on modelling energy adjusted variables.

| | | Quintile of energy adjusted EPA intake (gm) | | | P _{Trend} |
|---------------------------|------------------------|---|------------------------|------------------------|--------------------|
| | | Q1 0.01 [0.01 0.01] | Q3 0.04 [0.04 0.05] | Q5 0.12 [0.11 0.16] | |
| LDL-C (mg/dL) | Raw mean | 123 ± 34 | 125 ± 34 | 124 ± 35 | |
| | Adj. mean ¹ | 123 [122 124] | 125 [124 126] | 124 [123 125] | 0.29 |
| | Adj. mean ² | 123 [122 125] | 124 [123 125] | 123 [122 125] | 0.50 |
| TC (mg/dL) | Raw mean | 210 ± 41 | 213 ± 42 | 213 ± 42 | |
| | Adj. mean ¹ | 210 [208 211] | 213 [212 214] | 213 [212 214] | 0.002 |
| | Adj. mean ² | 211 [210 212] | 211 [210 212] | 211 [209 212] | 0.39 |
| ApoB100 (mg/dL) | Raw mean | 103 ± 28 | 104 ± 28 | 103 ± 28 | |
| | Adj. mean ¹ | 103 [102 103] | 105 [104 105] | 103 [103 104] | 0.68 |
| | Adj. mean ² | 103 [102 104] | 103 [102 104] | 103 [101 103.7] | 0.52 |
| LDL Size (Ø nm) | Geom. mean | 21.04 [21.02 21.06] | 21.05 [21.03 21.07] | 21.12 [21.11 21.14] | |
| | Adj. mean ¹ | 21.04 [21.02 21.06] | 21.05 [21.03 21.07] | 21.12 [21.1 21.13] | <.0001 |
| | Adj. mean ² | 21.1 [21.08 21.13] | 21.06 [21.03 21.08] | 21.07 [21.04 21.1] | 0.52 |
| LDL Particles | | | | | |
| Total (nm/L) | Geom. mean | 1189 [1179 1200] | 1211 [1198 1225] | 1190 [1180 1201] | |
| | Adj. mean ¹ | 1189 [1178 1199] | 1214 [1200 1227] | 1191 [1180 1202] | 0.41 |
| | Adj. mean ² | 1184 [1169 1199] | 1197 [1183 1210] | 1182 [1165 1199] | 0.30 |
| IDL (nm/L) | Geom. mean | 146 [143 148] | 145 [141 148] | 147 [145 150] | |
| | Adj. mean ¹ | 146 [143 148] | 145 [142 149] | 146 [143 149] | 0.96 |
| | Adj. mean ² | 146 [142 151] | 145 [141 149] | 144 [139 149] | 0.68 |
| Large (nm/L) | Geom. mean | 463 [454 473] | 481 [469 494] | 515 [505 525] | |
| | Adj. mean ¹ | 462 [453 471] | 482 [470 494] | 512 [501 523] | <.0001 |
| | Adj. mean ² | 491 [475 507] | 480 [467 494] | 488 [470 506] | 0.89 |
| Small A (nm/L) | Geom. mean | 62.2 [61.3 63.2] | 64 [62.6 65.5] | 65.8 [64.4 67.2] | |
| | Adj. mean ¹ | 62.3 [61.3 63.2] | 64 [62.6 65.4] | 66 [64.5 67.4] | <.001 |
| | Adj. mean ² | 62.4 [60.7 64.2] | 63.9 [62.3 65.6] | 65.7 [63.2 68.2] | 0.16 |
| Small B (nm/L) | Geom. mean | 661 [650 671] | 662 [648 676] | 647 [635 658] | |
| | Adj. mean ¹ | 662 [651 673] | 663 [649 676] | 644 [632 656] | 0.02 |
| | Adj. mean ² | 657 [640 673] | 654 [640 669] | 637 [619 656] | 0.27 |

Raw means ± sd and geometric (Geom.) means [CI]. ¹ Model 1: The adjusted means (Adj.) are estimated from linear regression models adjusted for age (continuous), total energy (quintiles), energy adjusted saturated fats (quintiles), energy adjusted monounsaturated fats (quintiles), energy adjusted *trans*-fat (quintiles), energy adjusted total n-6 (quintiles) and energy adjusted proteins (quintiles). *Is-means.

² Model 2: Adjusted as in model 1 but including smoking (current, past, never), alcohol use (rarely/never, 1–3 drinks/month, 1–6 drinks/week, and 1 drinks/day), BMI (continuous), exercise (quintiles of metabolic equivalent task hours per week), menopausal status (premenopausal, uncertain, postmenopausal), use of HT (current, past/never), hypertension (systolic blood pressure of at least 140 mmHg, diastolic blood pressure of at least 90 mmHg), antihypertensive treatment (yes or no), hypercholesterolemia (total cholesterol of at least 240 mg/dL), treatment for high cholesterol (yes or no), parental history of CHD (yes or no), energy adjusted glycemic index (quintiles), multivitamin use (current, past and never), aspirin use (Current use > 1x/week), red meat consumption, fruits and vegetables consumption (both quintiles) and energy adjusted ALA and DHA fatty acids (both in quintiles).

Table s5- LDL related variables. Strategy based on modelling energy adjusted variables.

| | | Quintile of energy adjusted DHA intake (gm) | | | |
|----------------------|------------------------|---|---------------------|---------------------|--------------------|
| | | Q2 | Q3 | Q5 | P _{Trend} |
| | | 0.09 [0.08 0.1] | 0.12 [0.11 0.13] | 0.28 [0.24 0.35] | |
| LDL-C | Raw mean | 124 ± 33 | 125 ± 35 | 125 ± 35 | |
| (mg/dL) | Adj. mean ¹ | 124 [123 125] | 125 [124 126] | 125 [124 126] | 0.005 |
| | Adj. mean ² | 123 [122 124] | 124 [123 125] | 124 [123 126] | 0.23 |
| TC | Raw mean | 212 ± 41 | 212 ± 42 | 214 ± 42 | |
| (mg/dL) | Adj. mean ¹ | 212 [210 213] | 213 [211 214] | 214 [213 215] | <.0001 |
| | Adj. mean ² | 211 [210 212] | 212 [211 213] | 212 [211 214] | 0.11 |
| ApoB100 | Raw mean | 104 ± 28 | 104 ± 28 | 104 ± 28 | |
| (mg/dL) | Adj. mean ¹ | 104 [103 104] | 104 [103 104] | 104 [103 105] | 0.01 |
| | Adj. mean ² | 103 [102 104] | 103 [102 103] | 104 [102 104.8] | 0.27 |
| LDL Size | Geom. mean | 21.06 [21.04 21.08] | 21.08 [21.06 21.1] | 21.11 [21.09 21.13] | |
| (Ø nm) | Adj. mean ¹ | 21.06 [21.04 21.08] | 21.08 [21.06 21.1] | 21.11 [21.09 21.13] | <.0001 |
| | Adj. mean ² | 21.07 [21.04 21.09] | 21.09 [21.07 21.11] | 21.12 [21.08 21.15] | 0.0075 |
| LDL Particles | | | | | |
| Total | Geom. mean | 1198 [1187 1210] | 1201 [1190 1212] | 1203 [1192 1214] | |
| (nm/L) | Adj. mean ¹ | 1197 [1186 1208] | 1201 [1190 1211] | 1204 [1192 1216] | 0.09 |
| | Adj. mean ² | 1186 [1173 1199] | 1187 [1176 1198] | 1198 [1180 1217] | 0.38 |
| IDL | Geom. mean | 147 [144 150] | 146 [144 149] | 147 [144 150] | |
| (nm/L) | Adj. mean ¹ | 147 [144 150] | 147 [144 150] | 146 [143 149] | 0.85 |
| | Adj. mean ² | 146 [142 150] | 145 [142 149] | 146 [141 152] | 0.83 |
| Large | Geom. mean | 476 [466 487] | 489 [479 499] | 507 [496 518] | |
| (nm/L) | Adj. mean ¹ | 477 [467 487] | 490 [480 500] | 508 [497 519] | <.0001 |
| | Adj. mean ² | 479 [465 492] | 493 [482 505] | 504 [485 524] | 0.02 |
| Small A | Geom. mean | 63.7 [62.5 64.9] | 63.3 [61.9 64.7] | 65.3 [63.9 66.7] | |
| (nm/L) | Adj. mean ¹ | 63.5 [62.3 64.8] | 63.2 [61.9 64.6] | 65.4 [63.9 66.9] | 0.005 |
| | Adj. mean ² | 64.2 [62.6 66] | 62.9 [61.4 64.5] | 63.3 [60.9 65.9] | 0.72 |
| Small B | Geom. mean | 655 [644 667] | 651 [640 663] | 658 [647 670] | |
| (nm/L) | Adj. mean ¹ | 657 [645 668] | 651 [641 663] | 655 [643 667] | 0.23 |
| | Adj. mean ² | 650 [636 664] | 648 [636 660] | 654 [635 674] | 0.96 |

Raw means ± sd and geometric (Geom.) means [CI]. ¹ Model 1: The adjusted means (Adj.) are estimated from linear regression models adjusted for age (continuous), total energy (quintiles), energy adjusted saturated fats (quintiles), energy adjusted monounsaturated fats (quintiles), energy adjusted *trans* fat (quintiles), energy adjusted total n-6 (quintiles) and energy adjusted proteins (quintiles).

² Model 2: Adjusted as in model 1 but including smoking (current, past, never), alcohol use (rarely/never, 1–3 drinks/month, 1–6 drinks/week, and 1 drinks/day), BMI (continuous), exercise (quintiles of metabolic equivalent task hours per week), menopausal status (premenopausal, uncertain, postmenopausal), use of HT (current, past/never), hypertension (systolic blood pressure of at least 140 mmHg, diastolic blood pressure of at least 90 mmHg), antihypertensive treatment (yes or no), hypercholesterolemia (total cholesterol of at least 240 mg/dL), treatment for high cholesterol (yes or no), parental history of CHD (yes or no), energy adjusted glycemic index (quintiles), multivitamin use (current, past and never), aspirin use (Current use > 1x/week), red meat consumption, fruits and vegetables consumption (both quintiles) and energy adjusted ALA and EPA fatty acids (both in quintiles).

Table s6- LDL related variables. Strategy based on modelling energy adjusted variables.

| | | Quintile of energy adjusted ALA intake (gm) | | | |
|----------------------|------------------------|---|---------------------|---------------------|--------------------|
| | | Q1 | Q3 | Q5 | P _{Trend} |
| | | 0.79 [0.71 0.84] | 1.12 [1.08 1.16] | 1.61 [1.5 1.79] | |
| LDL-C | Raw mean | 123 ± 34 | 124 ± 35 | 125 ± 34 | |
| (mg/dL) | Adj. mean ¹ | 125 [124 126] | 124 [123 125] | 124 [122 125] | 0.10 |
| | Adj. mean ² | 125 [124 126] | 123 [123 124] | 123 [122 124] | 0.03 |
| TC | Raw mean | 211 ± 42 | 212 ± 42 | 213 ± 42 | |
| (mg/dL) | Adj. mean ¹ | 213 [211 214] | 212 [211 213] | 211 [210 213] | 0.38 |
| | Adj. mean ² | 212 [211 213] | 211 [210 212] | 211 [210 212] | 0.25 |
| ApoB100 | Raw mean | 103 ± 28 | 104 ± 28 | 104 ± 28 | |
| (mg/dL) | Adj. mean ¹ | 104 [103 105] | 104 [103 104] | 103 [102 104] | 0.19 |
| | Adj. mean ² | 104 [103 105] | 103 [102 104] | 103 [102 103.4] | 0.09 |
| LDL Size | Geom. mean | 21.08 [21.06 21.1] | 21.07 [21.06 21.09] | 21.05 [21.04 21.07] | |
| (Ø nm) | Adj. mean ¹ | 21.04 [21.02 21.07] | 21.08 [21.06 21.1] | 21.08 [21.06 21.11] | 0.03 |
| | Adj. mean ² | 21.05 [21.03 21.07] | 21.08 [21.07 21.1] | 21.1 [21.07 21.12] | 0.002 |
| LDL Particles | | | | | |
| Total | Geom. mean | 1189 [1178 1199] | 1200 [1189 1211] | 1212 [1202 1223] | |
| (nm/L) | Adj. mean ¹ | 1208 [1196 1221] | 1199 [1188 1209] | 1195 [1181 1208] | 0.23 |
| | Adj. mean ² | 1203 [1191 1215] | 1189 [1180 1199] | 1186 [1173 1199] | 0.12 |
| IDL | Geom. mean | 152 [149 155] | 146 [144 149] | 141 [138 144] | |
| (nm/L) | Adj. mean ¹ | 147 [144 151] | 146 [144 149] | 146 [142 149] | 0.80 |
| | Adj. mean ² | 146 [143 150] | 145 [143 148] | 145 [142 149] | 0.95 |
| Large | Geom. mean | 474 [464 484] | 490 [480 500] | 493 [483 503] | |
| (nm/L) | Adj. mean ¹ | 472 [460 484] | 492 [482 503] | 489 [477 502] | 0.09 |
| | Adj. mean ² | 476 [463 489] | 493 [482 503] | 491 [477 505] | 0.15 |
| Small A | Geom. mean | 60.7 [59.4 62.1] | 64.1 [62.9 65.4] | 65.8 [64.4 67.1] | |
| (nm/L) | Adj. mean ¹ | 63.1 [61.6 64.7] | 64 [62.7 65.3] | 63.4 [61.7 65.1] | 0.84 |
| | Adj. mean ² | 62.5 [60.8 64.2] | 63.8 [62.4 65.1] | 63.2 [61.5 65.1] | 0.98 |
| Small B | Geom. mean | 654 [643 665] | 652 [641 663] | 669 [658 681] | |
| (nm/L) | Adj. mean ¹ | 663 [650 677] | 652 [641 663] | 659 [645 673] | 0.78 |
| | Adj. mean ² | 661 [648 675] | 648 [637 659] | 655 [641 669] | 0.80 |

Raw means ± sd and geometric (Geom.) means [CI]. ¹ Model 1: The adjusted means (Adj.) are estimated from linear regression models adjusted for age (continuous), total energy (quintiles), energy adjusted saturated fats (quintiles), energy adjusted monounsaturated fats (quintiles), energy adjusted *trans* fat (quintiles), energy adjusted total n-6 (quintiles) and energy adjusted proteins (quintiles).

² Model 2: Adjusted as in model 1 but including smoking (current, past, never), alcohol use (rarely/never, 1–3 drinks/month, 1–6 drinks/week, and 1 drinks/day), BMI (continuous), exercise (quintiles of metabolic equivalent task hours per week), menopausal status (premenopausal, uncertain, postmenopausal), use of HT (current, past/never), hypertension (systolic blood pressure of at least 140 mmHg, diastolic blood pressure of at least 90 mmHg), antihypertensive treatment (yes or no), hypercholesterolemia (total cholesterol of at least 240 mg/dL), treatment for high cholesterol (yes or no), parental history of CHD (yes or no), energy adjusted glycemic index (quintiles), multivitamin use (current, past and never), aspirin use (Current use > 1x/week), red meat consumption, fruits and vegetables consumption (both quintiles) and energy adjusted EPA and DHA fatty acids (both in quintiles).

Table s7- VLDL related variables. Strategy based on modelling energy adjusted variables.

| | | Quintile of energy adjusted EPA intake (gm) | | | |
|-----------------------------------|------------------------|---|---------------------|---------------------|--------------------|
| | | Q1 | Q3 | Q5 | P _{Trend} |
| | | 0.01 [0.01 0.01] | 0.04 [0.04 0.05] | 0.12 [0.11 0.16] | |
| TG (mg/dL) | Geom. mean | 124 [122 125] | 124 [122 127] | 117 [115 119] | |
| | Adj. mean ¹ | 124 [122 126] | 125 [122 127] | 117 [115 119] | <.0001 |
| | Adj. mean ² | 121 [118 124] | 123 [120 125] | 118 [115 121] | 0.19 |
| VLDL Size (Ø nm) | Geom. mean | 51.11 [50.92 51.31] | 51.05 [50.81 51.3] | 50.87 [50.68 51.07] | |
| | Adj. mean ¹ | 51.12 [50.93 51.31] | 51.1 [50.86 51.35] | 50.87 [50.67 51.07] | 0.07 |
| | Adj. mean ² | 50.93 [50.62 51.24] | 51.06 [50.79 51.32] | 50.91 [50.58 51.25] | 0.87 |
| VLDL Particles | | | | | |
| Total (nm/L) | Geom. mean | 57.3 [56.7 58] | 56.7 [55.8 57.7] | 53.1 [52.4 53.8] | |
| | Adj. mean ¹ | 57.3 [56.6 58] | 56.6 [55.8 57.5] | 53.4 [52.7 54.1] | <.0001 |
| | Adj. mean ² | 55.9 [54.8 57.1] | 56.4 [55.4 57.4] | 54.1 [53 55.4] | 0.03 |
| Large (nm/L) | Geom. mean | 2.62 [2.54 2.69] | 2.53 [2.45 2.63] | 2.21 [2.14 2.28] | |
| | Adj. mean ¹ | 2.61 [2.54 2.69] | 2.56 [2.47 2.66] | 2.23 [2.16 2.29] | <.0001 |
| | Adj. mean ² | 2.46 [2.35 2.58] | 2.51 [2.41 2.6] | 2.3 [2.19 2.41] | 0.07 |
| Medium (nm/L) | Geom. mean | 13.4 [13.1 13.7] | 13.2 [12.9 13.6] | 12.2 [11.9 12.5] | |
| | Adj. mean ¹ | 13.5 [13.2 13.8] | 13.2 [12.9 13.6] | 12.1 [11.8 12.4] | <.0001 |
| | Adj. mean ² | 12.7 [12.3 13.2] | 13.1 [12.6 13.5] | 12.5 [12 13.1] | 0.26 |
| Small (nm/L) | Geom. mean | 36.8 [36.2 37.3] | 36.4 [35.7 37.1] | 34.2 [33.6 34.8] | |
| | Adj. mean ¹ | 36.5 [36 37.1] | 36.3 [35.6 36.9] | 34.6 [34 35.1] | <.0001 |
| | Adj. mean ² | 36.1 [35.2 37] | 36.2 [35.5 37] | 34.8 [33.8 35.7] | 0.07 |
| VLDL-TG (mg/dL) | Geom. mean | 76.5 [75.5 77.6] | 75.6 [74.3 76.9] | 70.3 [69.3 71.3] | |
| | Adj. mean ¹ | 76.6 [75.6 77.6] | 75.7 [74.4 77] | 70.6 [69.6 71.6] | <.0001 |
| | Adj. mean ² | 74.2 [72.7 75.9] | 75 [73.7 76.4] | 71.6 [70 73.3] | 0.02 |

Raw means \pm sd and geometric (Geom.) means [CI]. ¹ Model 1: The adjusted means (Adj.) are estimated from linear regression models adjusted for age (continuous), total energy (quintiles), energy adjusted saturated fats (quintiles), energy adjusted monounsaturated fats (quintiles), energy adjusted *trans*-fat (quintiles), energy adjusted total n-6 (quintiles) and energy adjusted proteins (quintiles).

² Model 2: Adjusted as in model 1 but including smoking (current, past, never), alcohol use (rarely/never, 1–3 drinks/month, 1–6 drinks/week, and 1 drinks/day), BMI (continuous), exercise (quintiles of metabolic equivalent task hours per week), menopausal status (premenopausal, uncertain, postmenopausal), use of HT (current, past/never), hypertension (systolic blood pressure of at least 140 mmHg, diastolic blood pressure of at least 90 mmHg), antihypertensive treatment (yes or no), hypercholesterolemia (total cholesterol of at least 240 mg/dL), treatment for high cholesterol (yes or no), parental history of CHD (yes or no), energy adjusted glycemic index (quintiles), multivitamin use (current, past and never), aspirin use (Current use > 1x/week), red meat consumption, fruits and vegetables consumption (both quintiles) and energy adjusted ALA and DHA fatty acids (both in quintiles).

Table s8- VLDL related variables. Strategy based on modelling energy adjusted variables.

| | | Quintile of energy adjusted DHA intake (gm) | | | |
|---|------------------------|---|---------------------|---------------------|--------------------|
| | | Q1 | Q3 | Q5 | P ^{Trend} |
| | | 0.05 [0.03 0.06] | 0.12 [0.11 0.13] | 0.28 [0.24 0.35] | |
| TG (mg/dL) | Geom. mean | 123 [121 125] | 121 [119 123] | 119 [117 121] | |
| | Adj. mean ¹ | 123 [122 125] | 121 [119 123] | 118 [117 120] | <.0001 |
| | Adj. mean ² | 122 [119 125] | 119 [117 121] | 119 [116 122] | 0.35 |
| VLDL Size (Ø nm) | Geom. mean | 51.18 [51 51.36] | 50.77 [50.58 50.97] | 50.92 [50.72 51.12] | |
| | Adj. mean ¹ | 51.17 [50.99 51.36] | 50.78 [50.59 50.98] | 50.91 [50.7 51.12] | 0.30 |
| | Adj. mean ² | 51.22 [50.92 51.53] | 50.77 [50.56 50.99] | 50.83 [50.47 51.19] | 0.39 |
| VLDL Particles | | | | | |
| Total (nm/L) | Geom. mean | 57.2 [56.5 57.8] | 56.3 [55.6 57.1] | 53.9 [53.1 54.6] | |
| | Adj. mean ¹ | 57.2 [56.5 57.9] | 56.2 [55.5 56.9] | 54 [53.2 54.7] | <.0001 |
| | Adj. mean ² | 56.3 [55.2 57.4] | 55.3 [54.5 56.1] | 55.4 [54.1 56.8] | 0.61 |
| Large (nm/L) | Geom. mean | 2.63 [2.57 2.7] | 2.43 [2.36 2.5] | 2.27 [2.2 2.34] | |
| | Adj. mean ¹ | 2.63 [2.56 2.7] | 2.43 [2.36 2.5] | 2.28 [2.21 2.35] | <.0001 |
| | Adj. mean ² | 2.58 [2.47 2.7] | 2.36 [2.29 2.44] | 2.35 [2.23 2.48] | 0.12 |
| Medium (nm/L) | Geom. mean | 13.4 [13.2 13.7] | 13.2 [12.9 13.5] | 12.3 [12 12.7] | |
| | Adj. mean ¹ | 13.6 [13.3 13.9] | 13.2 [12.8 13.5] | 12.2 [11.9 12.5] | <.0001 |
| | Adj. mean ² | 13.5 [13 14] | 12.7 [12.4 13.1] | 12.5 [12 13.1] | 0.10 |
| Small (nm/L) | Geom. mean | 36.5 [36 37] | 36.1 [36.5 36.7] | 34.6 [34.1 35.2] | |
| | Adj. mean ¹ | 36.4 [35.8 36.9] | 36 [35.5 36.6] | 34.9 [34.4 35.5] | <.001 |
| | Adj. mean ² | 35.7 [34.8 36.6] | 35.6 [35 36.2] | 36 [34.9 37] | 0.70 |
| VLDL- TG (mg/dL) | Geom. mean | 76.5 [75.6 77.4] | 74.2 [73.2 75.2] | 71.4 [70.4 72.5] | |
| | Adj. mean ¹ | 76.7 [75.7 77.7] | 74.1 [73.1 75.1] | 71.4 [70.3 72.5] | <.0001 |
| | Adj. mean ² | 75.5 [73.9 77.1] | 72.7 [71.6 73.8] | 73.1 [71.3 75] | 0.30 |

Raw means \pm sd and geometric (Geom.) means [CI]. ¹ Model 1: The adjusted means (Adj.) are estimated from linear regression models adjusted for age (continuous), total energy (quintiles), energy adjusted saturated fats (quintiles), energy adjusted monounsaturated fats (quintiles), energy adjusted *trans* fat (quintiles), energy adjusted total n-6 (quintiles) and energy adjusted proteins (quintiles).

² Model 2: Adjusted as in model 1 but including smoking (current, past, never), alcohol use (rarely/never, 1–3 drinks/month, 1–6 drinks/week, and 1 drinks/day), BMI (continuous), exercise (quintiles of metabolic equivalent task hours per week), menopausal status (premenopausal, uncertain, postmenopausal), use of HT (current, past/never), hypertension (systolic blood pressure of at least 140 mmHg, diastolic blood pressure of at least 90 mmHg), antihypertensive treatment (yes or no), hypercholesterolemia (total cholesterol of at least 240 mg/dL), treatment for high cholesterol (yes or no), parental history of CHD (yes or no), energy adjusted glycemic index (quintiles), multivitamin use (current, past and never), aspirin use (Current use > 1x/week), red meat consumption, fruits and vegetables consumption (both quintiles) and energy adjusted ALA and EPA fatty acids (both in quintiles).

Table s9- VLDL related variables. Strategy based on modelling energy adjusted variables.

| | | Quintile of energy adjusted ALA intake (gm) | | | P _{Trend} |
|-----------------------------------|------------------------|---|---------------------|---------------------|--------------------|
| | | Q1 | Q3 | Q5 | |
| | | 0.79 [0.71 0.84] | 1.12 [1.08 1.16] | 1.61 [1.5 1.79] | |
| TG (mg/dL) | Geom. mean | 122 [120 123] | 122 [121 124] | 122 [121 124] | |
| | Adj. mean ¹ | 123 [121 125] | 122 [120 124] | 122 [119 124] | 0.62 |
| | Adj. mean ² | 122 [120 124] | 121 [119 123] | 121 [118 123] | 0.64 |
| VLDL Size (Ø nm) | Geom. mean | 51.53 [51.34 51.74] | 51 [50.81 51.19] | 50.61 [50.42 50.8] | |
| | Adj. mean ¹ | 51.26 [51.03 51.5] | 51.03 [50.83 51.22] | 50.84 [50.59 51.09] | 0.04 |
| | Adj. mean ² | 51.25 [51 51.5] | 50.98 [50.78 51.17] | 50.72 [50.46 50.97] | 0.01 |
| VLDL Particles | | | | | |
| Total (nm/L) | Geom. mean | 54.2 [53.5 54.9] | 56.6 [55.9 57.3] | 57.6 [56.9 58.3] | |
| | Adj. mean ¹ | 56.3 [55.4 57.2] | 56.4 [55.7 57.1] | 55.7 [54.8 56.6] | 0.45 |
| | Adj. mean ² | 55.7 [54.8 56.7] | 56.2 [55.5 56.9] | 55.8 [54.8 56.7] | 0.94 |
| Large (nm/L) | Geom. mean | 2.63 [2.56 2.7] | 2.5 [2.42 2.57] | 2.36 [2.29 2.43] | |
| | Adj. mean ¹ | 2.59 [2.5 2.68] | 2.49 [2.41 2.56] | 2.43 [2.34 2.52] | 0.06 |
| | Adj. mean ² | 2.56 [2.47 2.66] | 2.45 [2.38 2.52] | 2.38 [2.29 2.47] | 0.04 |
| Medium (nm/L) | Geom. mean | 12.9 [12.6 13.1] | 13.3 [13 13.7] | 13.1 [12.8 13.5] | |
| | Adj. mean ¹ | 13.2 [12.8 13.6] | 13.3 [12.9 13.6] | 13 [12.6 13.4] | 0.65 |
| | Adj. mean ² | 13 [12.6 13.4] | 13.1 [12.8 13.4] | 12.9 [12.5 13.3] | 0.88 |
| Small (nm/L) | Geom. mean | 34.1 [33.5 34.7] | 36.3 [35.7 36.8] | 37.3 [36.8 37.9] | |
| | Adj. mean ¹ | 35.7 [35 36.4] | 36.2 [35.6 36.7] | 35.8 [35.1 36.5] | 0.96 |
| | Adj. mean ² | 35.4 [34.7 36.2] | 36.1 [35.5 36.7] | 35.9 [35.2 36.7] | 0.61 |
| VLDL-TG (mg/dL) | Geom. mean | 74.1 [73.1 75.1] | 75.3 [74.3 76.3] | 74.9 [73.9 76] | |
| | Adj. mean ¹ | 75.6 [74.4 76.9] | 75.1 [74.1 76.1] | 73.8 [72.5 75.1] | 0.11 |
| | Adj. mean ² | 74.8 [73.5 76.1] | 74.6 [73.5 75.6] | 73.4 [72.1 74.7] | 0.23 |

Raw means ± sd and geometric (Geom.) means [CI]. ¹ Model 1: The adjusted means (Adj.) are estimated from linear regression models adjusted for age (continuous), total energy (quintiles), energy adjusted saturated fats (quintiles), energy adjusted monounsaturated fats (quintiles), energy adjusted *trans* fat (quintiles), energy adjusted total n-6 (quintiles) and energy adjusted proteins (quintiles).

² Model 2: Adjusted as in model 1 but including smoking (current, past, never), alcohol use (rarely/never, 1–3 drinks/month, 1–6 drinks/week, and 1 drinks/day), BMI (continuous), exercise (quintiles of metabolic equivalent task hours per week), menopausal status (premenopausal, uncertain, postmenopausal), use of HT (current, past/never), hypertension (systolic blood pressure of at least 140 mmHg, diastolic blood pressure of at least 90 mmHg), antihypertensive treatment (yes or no), hypercholesterolemia (total cholesterol of at least 240 mg/dL), treatment for high cholesterol (yes or no), parental history of CHD (yes or no), energy adjusted glycemic index (quintiles), multivitamin use (current, past and never), aspirin use (Current use > 1x/week), red meat consumption, fruits and vegetables consumption (both quintiles) and energy adjusted EPA and DHA fatty acids (both in quintiles).

Table s10- HDL related variables. Strategy based on modelling energy adjusted variables.

| | | Quintile of energy adjusted EPA intake (gm) | | | P _{Trend} |
|----------------------|------------------------|---|------------------|------------------|--------------------|
| | | Q1 | Q3 | Q5 | |
| | | 0.01 [0.01 0.01] | 0.04 [0.04 0.05] | 0.12 [0.11 0.16] | |
| HDL-C | Raw mean | 52.9 ± 14.5 | 53.8 ± 15 | 55.9 ± 15.7 | |
| (mg/dL) | Adj. mean ¹ | 52.9 [52.5 53.3] | 53.8 [53.3 54.3] | 55.5 [55.1 55.9] | <.0001 |
| | Adj. mean ² | 54.4 [53.8 55] | 53.8 [53.3 54.4] | 53.9 [53.3 54.6] | 0.30 |
| ApoA1 | Raw mean | 149 ± 25 | 152 ± 25 | 154 ± 26 | |
| (mg/dL) | Adj. mean ¹ | 149 [149 150] | 152 [151 152] | 153 [152 154] | <.0001 |
| | Adj. mean ² | 152 [151 153] | 151 [150 152] | 150 [149 151] | 0.02 |
| HDL Size | Geom. mean | 9.17 [9.16 9.19] | 9.18 [9.16 9.2] | 9.25 [9.24 9.27] | |
| (Ø nm) | Adj. mean ¹ | 9.18 [9.16 9.19] | 9.18 [9.16 9.2] | 9.24 [9.23 9.26] | <.0001 |
| | Adj. mean ² | 9.23 [9.21 9.25] | 9.19 [9.17 9.2] | 9.2 [9.18 9.22] | 0.55 |
| HDL Particles | | | | | |
| Total | Geom. mean | 36.6 [36.4 36.8] | 37.1 [36.9 37.4] | 37.2 [37 37.4] | |
| (µm/L) | Adj. mean ¹ | 36.7 [36.5 36.9] | 37.1 [36.9 37.3] | 37 [36.8 37.2] | 0.44 |
| | Adj. mean ² | 37 [36.7 37.3] | 37 [36.7 37.2] | 36.5 [36.2 36.8] | 0.004 |
| Large | Geom. mean | 5.32 [5.24 5.4] | 5.42 [5.32 5.53] | 5.86 [5.77 5.96] | |
| (µm/L) | Adj. mean ¹ | 5.35 [5.27 5.43] | 5.41 [5.3 5.51] | 5.76 [5.67 5.86] | <.0001 |
| | Adj. mean ² | 5.66 [5.53 5.8] | 5.44 [5.33 5.55] | 5.47 [5.33 5.61] | 0.16 |
| Medium | Geom. mean | 11.3 [11.1 11.4] | 11.4 [11.2 11.6] | 11.3 [11.1 11.5] | |
| (µm/L) | Adj. mean ¹ | 11.5 [11.3 11.6] | 11.4 [11.2 11.6] | 11.1 [10.9 11.3] | 0.01 |
| | Adj. mean ² | 11.5 [11.2 11.8] | 11.3 [11.1 11.6] | 10.9 [10.7 11.2] | 0.01 |
| Small | Geom. mean | 17.6 [17.5 17.8] | 17.8 [17.6 18] | 17.5 [17.4 17.7] | |
| (µm/L) | Adj. mean ¹ | 17.5 [17.4 17.7] | 17.8 [17.7 18] | 17.7 [17.5 17.9] | 0.70 |
| | Adj. mean ² | 17.3 [17.1 17.6] | 17.7 [17.5 17.9] | 17.8 [17.5 18] | 0.54 |

Raw means ± sd and geometric (Geom.) means [CI]. ¹ Model 1: The adjusted means (Adj.) are estimated from linear regression models adjusted for age (continuous), total energy (quintiles), energy adjusted saturated fats (quintiles), energy adjusted monounsaturated fats (quintiles), energy adjusted *trans* fat (quintiles), energy adjusted total n-6 (quintiles) and energy adjusted proteins (quintiles).

² Model 2: Adjusted as in model 1 but including smoking (current, past, never), alcohol use (rarely/never, 1–3 drinks/month, 1–6 drinks/week, and 1 drinks/day), BMI (continuous), exercise (quintiles of metabolic equivalent task hours per week), menopausal status (premenopausal, uncertain, postmenopausal), use of HT (current, past/never), hypertension (systolic blood pressure of at least 140 mmHg, diastolic blood pressure of at least 90 mmHg), antihypertensive treatment (yes or no), hypercholesterolemia (total cholesterol of at least 240 mg/dL), treatment for high cholesterol (yes or no), parental history of CHD (yes or no), energy adjusted glycemic index (quintiles), multivitamin use (current, past and never), aspirin use (Current use > 1x/week), red meat consumption, fruits and vegetables consumption (both quintiles) and energy adjusted ALA and DHA fatty acids (both in quintiles)

Table s11- HDL related variables. Strategy based on modelling energy adjusted variables.

| | | Quintile of energy adjusted DHA intake (gm) | | | |
|----------------------------------|------------------------|---|------------------|------------------|---------------------|
| | | Q1 | Q3 | Q5 | P _{linear} |
| | | 0.05 [0.03 0.06] | 0.12 [0.11 0.13] | 0.28 [0.24 0.35] | |
| HDL-C (mg/dL) | Raw mean | 53 ± 14.6 | 54.2 ± 14.9 | 55.3 ± 15.3 | |
| | Adj. mean ¹ | 53.1 [52.7 53.5] | 54.3 [53.8 54.7] | 55.1 [54.7 55.6] | <.0001 |
| | Adj. mean ² | 53.3 [52.7 54] | 54.6 [54.2 55.1] | 55 [54.3 55.7] | 0.02 |
| ApoA1 (mg/dL) | Raw mean | 150 ± 25 | 152 ± 25 | 153 ± 26 | |
| | Adj. mean ¹ | 150 [149 150] | 152 [151 152] | 153 [152 154] | <.0001 |
| | Adj. mean ² | 150 [149 151] | 152 [151 153] | 153 [152 154] | 0.004 |
| HDL Size (Ø nm) | Geom. mean | 9.17 [9.16 9.18] | 9.19 [9.17 9.2] | 9.23 [9.22 9.25] | |
| | Adj. mean ¹ | 9.18 [9.16 9.19] | 9.19 [9.18 9.2] | 9.22 [9.21 9.24] | <.0001 |
| | Adj. mean ² | 9.17 [9.15 9.19] | 9.21 [9.19 9.22] | 9.24 [9.21 9.26] | 0.002 |
| HDL Particles | | | | | |
| Total (µm/L) | Geom. mean | 36.7 [36.6 36.9] | 37 [36.8 37.2] | 37.1 [36.9 37.3] | |
| | Adj. mean ¹ | 36.9 [36.7 37] | 37 [36.8 37.2] | 36.9 [36.7 37.1] | 0.58 |
| | Adj. mean ² | 36.9 [36.6 37.2] | 37 [36.8 37.1] | 37.1 [36.8 37.4] | 0.29 |
| Large (µm/L) | Geom. mean | 5.32 [5.25 5.4] | 5.48 [5.39 5.57] | 5.72 [5.62 5.82] | |
| | Adj. mean ¹ | 5.36 [5.28 5.44] | 5.48 [5.4 5.57] | 5.65 [5.55 5.74] | <.0001 |
| | Adj. mean ² | 5.33 [5.2 5.45] | 5.57 [5.48 5.66] | 5.74 [5.58 5.9] | 0.002 |
| Medium (µm/L) | Geom. mean | 11.3 [11.2 11.5] | 11.3 [11.1 11.5] | 11.3 [11.1 11.5] | |
| | Adj. mean ¹ | 11.5 [11.4 11.7] | 11.3 [11.1 11.5] | 11.1 [10.9 11.2] | 0.005 |
| | Adj. mean ² | 11.4 [11.2 11.7] | 11.3 [11.1 11.4] | 11.3 [11 11.6] | 0.94 |
| Small (µm/L) | Geom. mean | 17.7 [17.6 17.9] | 17.8 [17.7 18] | 17.6 [17.4 17.7] | |
| | Adj. mean ¹ | 17.6 [17.5 17.7] | 17.8 [17.6 18] | 17.7 [17.6 17.9] | 0.53 |
| | Adj. mean ² | 17.7 [17.5 18] | 17.7 [17.5 17.9] | 17.5 [17.2 17.8] | 0.33 |

Raw means ± sd and geometric (Geom.) means [CI]. ¹ Model 1: The adjusted means (Adj.) are estimated from linear regression models adjusted for age (continuous), total energy (quintiles), energy adjusted saturated fats (quintiles), energy adjusted monounsaturated fats (quintiles), energy adjusted *trans* fat (quintiles), energy adjusted total n-6 (quintiles) and energy adjusted proteins (quintiles).

² Model 2: Adjusted as in model 1 but including smoking (current, past, never), alcohol use (rarely/never, 1–3 drinks/month, 1–6 drinks/week, and 1 drinks/day), BMI (continuous), exercise (quintiles of metabolic equivalent task hours per week), menopausal status (premenopausal, uncertain, postmenopausal), use of HT (current, past/never), hypertension (systolic blood pressure of at least 140 mmHg, diastolic blood pressure of at least 90 mmHg), antihypertensive treatment (yes or no), hypercholesterolemia (total cholesterol of at least 240 mg/dL), treatment for high cholesterol (yes or no), parental history of CHD (yes or no), energy adjusted glycemic index (quintiles), multivitamin use (current, past and never), aspirin use (Current use > 1x/week), red meat consumption, fruits and vegetables consumption (both quintiles) and energy adjusted ALA and EPA fatty acids (both in quintiles).

Table s12- HDL related variables. Strategy based on modelling energy adjusted variables.

| | | Quintile of energy adjusted ALA intake (gm) | | | |
|----------------------|------------------------|---|------------------|------------------|--------------------|
| | | Q1 | Q3 | Q5 | P _{Trend} |
| | | 0.79 [0.71 0.84] | 1.12 [1.08 1.16] | 1.61 [1.5 1.79] | |
| HDL-C | Raw mean | 53.5 ± 14.6 | 53.9 ± 15 | 54.3 ± 15.2 | |
| (mg/dL) | Adj. mean ¹ | 53.7 [53.2 54.2] | 54 [53.6 54.5] | 53.9 [53.4 54.4] | 0.83 |
| | Adj. mean ² | 53.8 [53.3 54.3] | 54 [53.6 54.4] | 54.1 [53.6 54.7] | 0.48 |
| ApoA1 | Raw mean | 150 ± 25 | 151 ± 25 | 152 ± 26 | |
| (mg/dL) | Adj. mean ¹ | 151 [150 151] | 151 [151 152] | 151 [150 152] | 0.51 |
| | Adj. mean ² | 151 [150 152] | 151 [151 152] | 151 [150 152] | 0.50 |
| HDL Size | Geom. mean | 9.2 [9.19 9.21] | 9.18 [9.17 9.2] | 9.19 [9.18 9.21] | |
| (Ø nm) | Adj. mean ¹ | 9.18 [9.16 9.19] | 9.19 [9.17 9.2] | 9.21 [9.19 9.22] | 0.08 |
| | Adj. mean ² | 9.18 [9.17 9.2] | 9.2 [9.18 9.21] | 9.22 [9.2 9.24] | 0.01 |
| HDL Particles | | | | | |
| Total | Geom. mean | 36.9 [36.7 37] | 37 [36.8 37.2] | 37 [36.8 37.1] | |
| (µm/L) | Adj. mean ¹ | 37 [36.8 37.3] | 37 [36.8 37.2] | 36.7 [36.5 37] | 0.10 |
| | Adj. mean ² | 37 [36.8 37.3] | 37 [36.8 37.2] | 36.7 [36.5 36.9] | 0.09 |
| Large | Geom. mean | 5.4 [5.32 5.49] | 5.44 [5.35 5.53] | 5.55 [5.46 5.64] | |
| (µm/L) | Adj. mean ¹ | 5.37 [5.27 5.48] | 5.46 [5.37 5.55] | 5.52 [5.41 5.64] | 0.19 |
| | Adj. mean ² | 5.41 [5.31 5.51] | 5.5 [5.42 5.58] | 5.59 [5.48 5.7] | 0.07 |
| Medium | Geom. mean | 11.6 [11.5 11.8] | 11.3 [11.1 11.4] | 11 [10.9 11.2] | |
| (µm/L) | Adj. mean ¹ | 11.5 [11.3 11.7] | 11.3 [11.1 11.5] | 11.2 [11 11.4] | 0.09 |
| | Adj. mean ² | 11.4 [11.2 11.6] | 11.3 [11.2 11.5] | 11.2 [11 11.4] | 0.31 |
| Small | Geom. mean | 17.3 [17.2 17.5] | 17.8 [17.7 18] | 17.9 [17.8 18.1] | |
| (µm/L) | Adj. mean ¹ | 17.7 [17.5 17.9] | 17.8 [17.7 18] | 17.6 [17.4 17.8] | 0.23 |
| | Adj. mean ² | 17.8 [17.6 18] | 17.7 [17.5 17.9] | 17.4 [17.2 17.6] | 0.03 |

Raw means ± sd and geometric (Geom.) means [CI]. ¹ Model 1: The adjusted means (Adj.) are estimated from linear regression models adjusted for age (continuous), total energy (quintiles), energy adjusted saturated fats (quintiles), energy adjusted monounsaturated fats (quintiles), energy adjusted *trans*-fat (quintiles), energy adjusted total n-6 (quintiles) and energy adjusted proteins (quintiles).

² Model 2: Adjusted as in model 1 but including smoking (current, past, never), alcohol use (rarely/never, 1–3 drinks/month, 1–6 drinks/week, and 1 drinks/day), BMI (continuous), exercise (quintiles of metabolic equivalent task hours per week), menopausal status (premenopausal, uncertain, postmenopausal), use of HT (current, past/never), hypertension (systolic blood pressure of at least 140 mmHg, diastolic blood pressure of at least 90 mmHg), antihypertensive treatment (yes or no), hypercholesterolemia (total cholesterol of at least 240 mg/dL), treatment for high cholesterol (yes or no), parental history of CHD (yes or no), energy adjusted glycemic index (quintiles), multivitamin use (current, past and never), aspirin use (Current use > 1x/week), red meat consumption, fruits and vegetables consumption (both quintiles) and energy adjusted EPA and DHA fatty acids (both in quintiles).

Table s13- Correspondence between absolute values and per 1% and per 5% differences.

| | | Absolute values | 1% differences | 5 % differences |
|----------------|---------|-----------------|----------------|-----------------|
| LDL-C | (mg/dL) | 123 | 1.23 | 6.17 |
| TC | (mg/dL) | 210 | 2.10 | 10.51 |
| ApoB100 | (mg/dL) | 103 | 1.03 | 5.16 |
| LDL Size | (Ø nm) | 21 | 0.21 | 1.05 |
| LDL Particles | | | | |
| Total | (nm/L) | 1188 | 11.88 | 59.38 |
| IDL | (nm/L) | 151 | 1.51 | 7.56 |
| Large | (nm/L) | 460 | 4.60 | 22.98 |
| Small A | (nm/L) | 60 | 0.06 | 3.00 |
| Small B | (nm/L) | 658 | 6.58 | 32.90 |
| TG | (mg/dL) | 123 | 1.23 | 6.15 |
| VLDL Size | (Ø nm) | 52 | 0.52 | 2.58 |
| VLDL Particles | | | | |
| Total | (nm/L) | 55 | 0.55 | 2.77 |
| Large | (nm/L) | 3 | 0.03 | 0.14 |
| Medium | (nm/L) | 13 | 0.13 | 0.65 |
| Small | (nm/L) | 35 | 0.35 | 1.75 |
| VLDL-TG | (mg/dL) | 75 | 0.75 | 3.77 |
| HDL-C | (mg/dL) | 53 | 0.53 | 2.66 |
| ApoA1 | (mg/dL) | 150 | 1.50 | 7.49 |
| HDL Size | (Ø nm) | 9 | 0.09 | 0.46 |
| HDL Particles | | | | |
| Total | (µm/L) | 37 | 0.37 | 1.84 |
| Large | (µm/L) | 5 | 0.05 | 0.27 |
| Medium | (µm/L) | 12 | 0.12 | 0.58 |
| Small | (µm/L) | 17 | 0.17 | 0.87 |

The absolute values are the raw mean values of the n-3 1st quintile intake.

Table s14- Adjusted means (95% CI) for the lower exposure quintiles (Q1) that showed a significant association (Ptrend<0.05).

| | Fish | n-3 | EPA | DHA | ALA |
|-----------------------------|---------------------|---------------------|------------------|---------------------|---------------------|
| TC (mg/dL) | 210 (210–211) | | | | |
| Apo B100 (mg/dL) | 102 (102–103) | | | | |
| LDL-C (mg/dL) | 123 (122–124) | | | | 125 (124–126) |
| LDL particles (nm/L) | 1189 (1173–1198) | | | | |
| Large LDL particles (nm/L) | 469 (460–479) | 458 (447–470) | | 479 (465–492) | |
| LDL size (nm) | 21.06 (21.04–21.08) | 21.03 (21.01–21.05) | | 21.07 (21.04–21.09) | 21.05 (21.03–21.07) |
| TG (mg/dL) | 123 (122–124) | 124 (123–125) | | | |
| VLDL-TG (mg/dL) | 75.6 (74.6–76.6) | 76.6 (75.4–77.9) | 74.2 (72.7–75.9) | | |
| Total VLDL particles (nm/L) | 56.2 (55.5–56.9) | 57 (56.2–57.9) | 55.9 (54.8–57.1) | | |
| Large VLDL particles (nm/L) | 2.62 (2.55–2.7) | 2.69 (2.6–2.78) | | | 2.56 (2.47–2.66) |
| HDL-C (mg/dL) | | | | 53.3 (52.7–54) | |
| Large HDL particles (µm/L) | | 5.37 (5.28–5.47) | | 5.33 (5.2–5.45) | |
| HDL size (nm) | | 9.17 (9.16–9.19) | | 9.17 (9.15–9.19); | 9.18 (9.17–9.2) |

The means were adjusted for all demographic, clinical, and dietary factors. Blank spaces indicate no significant association (fish: Q1, n=5839; Q5, n=5617; n-3: Q1, n=5248; Q5, n=5123; EPA: Q1, n=5370; Q5, n=4947; 6 DHA: Q1, n=6351; Q5, n=4797; ALA: Q1, n=5286; Q5, n=5097). ALA indicates a-linolenic acid; Apo, apolipoprotein; DHA, docosahexaenoic acid; EPA, eicosapentaenoic acid; TC, total cholesterol; LDL, low density lipoprotein; LDL-C, low-density lipoprotein cholesterol; TG, triglycerides; VLDL, very low-density lipoprotein; HDL, high-density lipoprotein; HDL-C, high-density lipoprotein cholesterol.