

Supplementary data

Table S1: Cross-sectional association between circulating fatty acids and dairy fat intake

Table S2: Cross-sectional association between circulating fatty acids and fish fat intake

Table S3: Association between the circulating fatty acids EPA and DHA and fish fat intake

Table S4: Association between fatty acid markers from plasma triglycerides (TG) and cardiovascular risk factors.

Table S5: Association between fatty acid markers from plasma phospholipids (PL) and cardiovascular risk factors.

Table S6: Association between dairy fat intake and cardiovascular risk factors

Table S7: Association between fish fat intake and cardiovascular risk factors

Table S8A-C: Papers investigating the association between dairy fat biomarkers and cardiovascular health (based on an illustrative search)

References

Table S1: Cross-sectional association between circulating fatty acids and dairy fat intake

Total dairy Fat (g/d)	Fatty acids											
	C14:0		C15:0		C17:0		Trans-C16:1(n-7)		Trans-C18:1(n-7)		CLA	
	β	p-value	β	p-value	β	p-value	β	p-value	β	p-value	β	p-value
Plasma TG												
Model 1	0.222	<0.001	0.257	<0.001	0.030	0.43	0.188	<0.001	0.261	<0.001	0.171	<0.001
Model 2	0.175	<0.001	0.250	<0.001	0.048	0.15	0.205	<0.001	0.248	<0.001	0.188	<0.001
Model 3	0.162	<0.001	0.231	<0.001	0.043	0.18	0.189	<0.001	0.229	<0.001	0.173	<0.001
Model 4	0.167	<0.001	0.230	<0.001	0.040	0.21	0.194	<0.001	0.230	<0.001	0.175	<0.001
Model 5	0.183	<0.001	0.240	<0.001	0.040	0.20	0.191	<0.001	0.218	<0.001	0.182	<0.001
Model 6	0.168	<0.001	0.230	<0.001	0.041	0.20	0.195	<0.001	0.229	<0.001	0.175	<0.001
Model 7	0.204	<0.001	0.231	<0.001	0.042	0.27	0.193	<0.001	0.236	<0.001	0.182	<0.001
Plasma PL												
Model 1	0.198	<0.001	0.192	<0.001	0.127	<0.001	0.085	0.02	0.232	<0.001	0.228	<0.001
Model 2	0.202	<0.001	0.189	<0.001	0.119	<0.001	0.118	<0.001	0.239	<0.001	0.242	<0.001
Model 3	0.170	<0.001	0.162	<0.001	0.092	0.003	0.105	0.001	0.219	<0.001	0.217	<0.001
Model 4	0.174	<0.001	0.164	<0.001	0.090	0.004	0.110	<0.001	0.214	<0.001	0.220	<0.001
Model 5	0.182	<0.001	0.165	<0.001	0.127	<0.001	0.101	0.001	0.197	<0.001	0.211	<0.001
Model 6	0.174	<0.001	0.164	<0.001	0.090	0.003	0.111	<0.001	0.214	<0.001	0.220	<0.001
Model 7	0.177	<0.001	0.166	<0.001	0.094	0.003	0.109	<0.001	0.214	<0.001	0.229	<0.001

Model 1: Crude model; **Model 2:** Adjusted for energy intake; **Model 3:** Adjusted for Model 2 + age, sex, education and smoking; **Model 4:** Adjusted for model 2 + age, sex and BMI; **Model 5:** Adjusted for model 4 + total carbohydrates and Fat; **Model 6:** Adjusted for Model 4 + fish fat intake; **Model 7:** Adjusted for model 4 + Total Triglycerides + Total Cholesterol

Dependent: Dairy fat intake, independent: Circulating fatty acids. Associations between fatty acid status and dairy fat intake are reported as standardized β 's.

Abbreviations: C14:0, Myristic acid; C15:0, Pentadecanoic acid; C17:0, Heptadecanoic acid; Trans-C16:1(n-7), Trans-Palmitoleic acid; Trans-C18:1(n-7), Vaccenic acid; CLA, Conjugated Linoleic acid; TG, Triglycerides; PL, Phospholipids; BMI, Body Mass Index

Table S2: Cross-sectional association between circulating fatty acids and fish fat intake

Total fish Fat (g/d)	Fatty acids											
	C14:0		C15:0		C17:0		Trans-C16:1(n-7)		Trans-C18:1(n-7)		CLA	
	β	p-value	β	p-value	β	p-value	β	p-value	β	p-value	β	p-value
Plasma TG												
Model 1	-0.094	0.01	-0.045	0.22	0.080	0.04	0.022	0.60	-0.018	0.61	-0.093	0.01
Model 2	-0.101	0.01	-0.045	0.21	0.082	0.03	0.024	0.57	-0.020	0.58	-0.091	0.01
Model 3	-0.118	0.001	-0.069	0.06	0.093	0.02	0.001	0.98	-0.045	0.23	-0.106	0.004
Model 4	-0.104	0.01	-0.062	0.09	0.075	0.05	0.017	0.69	-0.030	0.42	-0.097	0.01
Model 5	-0.095	0.01	-0.045	0.21	0.087	0.02	0.019	0.63	-0.032	0.38	-0.101	0.004
Model 6	-0.104	0.01	-0.062	0.10	0.076	0.05	0.022	0.61	-0.027	0.48	-0.098	0.01
Model 7	-0.068	0.09	-0.058	0.11	0.023	0.61	0.008	0.85	-0.019	0.61	-0.082	0.02
Plasma PL												
Model 1	-0.064	0.07	-0.008	0.83	0.067	0.07	0.034	0.35	-0.065	0.89	-0.070	0.05
Model 2	-0.064	0.08	-0.008	0.83	0.066	0.07	0.039	0.29	-0.004	0.91	-0.068	0.06
Model 3	-0.087	0.02	-0.016	0.65	0.063	0.09	0.026	0.49	-0.024	0.53	-0.089	0.02
Model 4	-0.076	0.04	-0.020	0.59	0.049	0.19	0.034	0.36	-0.020	0.60	-0.078	0.03
Model 5	-0.068	0.06	-0.007	0.84	0.098	0.01	0.041	0.26	-0.014	0.72	-0.097	0.01
Model 6	-0.076	0.04	-0.017	0.65	0.051	0.17	0.037	0.33	-0.016	0.67	-0.079	0.04
Model 7	-0.077	0.04	-0.036	0.33	0.037	0.34	0.020	0.59	-0.029	0.44	-0.070	0.06

Model 1: Crude model; **Model 2:** Adjusted for energy intake; **Model 3:** Adjusted for Model 2 + age, sex, education and smoking; **Model 4:** Adjusted for Model 2 + age, sex and BMI; **Model 5:** Adjusted for model 4 + total carbohydrates and Fat; **Model 6:** Adjusted for Model 4 + dairy fat intake; **Model 7:** Adjusted for model 4 + Total Triglycerides + Total Cholesterol

Dependent: fish fat intake, independent: fatty acids. Associations between fatty acid status and fish fat intake are reported as standardized β 's. Abbreviations: C14:0, Myristic acid; C15:0, Pentadecanoic acid; C17:0, Heptadecanoic acid; Trans-C16:1(n-7), Trans-Palmitoleic acid; Trans-C18:1(n-7), Vaccenic acid; CLA, Conjugated Linoleic acid; TG, Triglycerides; PL, Phospholipids; BMI, Body Mass Index

Table S3: Association between the circulating fatty acids EPA and DHA and fish fat intake

Total fish Fat (g/d)	Omega-3 fatty acids			
	EPA		DHA	
	β	p-value	β	p-value
Plasma TG				
Model 1	0.277	<0.001	0.425	<0.001
Model 2	0.283	<0.001	0.440	<0.001
Model 3	0.272	<0.001	0.445	<0.001
Plasma PL				
Model 1	0.256	<0.001	0.457	<0.001
Model 2	0.258	<0.001	0.465	<0.001
Model 3	0.248	<0.001	0.464	<0.001

Model 1: Crude model; **Model 2:** Adjusted for model 1 + energy intake; **Model 3:** Adjusted for Model 2 + age and sex, education and smoking

Dependent: fish fat intake, independent: fatty acids. Associations between fatty acid status and fish fat intake are reported as standardized β 's.
Abbreviations: DHA, docosahexaenoic acid; EPA: eicosapentaenoic acid; PL, Phospholipids; TG, Triglycerides

Table S4: Association between fatty acid markers from plasma triglycerides (TG) and cardiovascular risk factors

	C14:0		C15:0		C17:0		Trans-C16:1(n-7)		Trans-C18:1(n-7)		CLA	
	Model S1	Model S2	Model S1	Model S2	Model S1	Model S2	Model S1	Model S2	Model S1	Model S2	Model S1	Model S2
Dependent variable												
Weight (kg)	0.074*	-0.025	-0.079*	-0.079	-0.116**	0.022	-0.032	-0.027	0.011	-0.026	0.000	-0.040
Waist circumference	0.094**	-0.041	-0.096**	-0.107***	-0.144***	0.039	-0.035	-0.040	0.023	-0.033	-0.007	-0.064*
BMI (kg/m ²)	0.081*	-0.066	-0.121**	-0.126***	-0.173***	0.022	0.078	-0.052	-0.010	-0.067*	-0.015	-0.075*
SBP (mmHg)	0.032	-0.047	-0.059	-0.066*	-0.074*	0.039	-0.024	-0.036	-0.013	-0.050	-0.044	-0.080
DBP (mmHg)	0.073*	0.001	-0.064*	-0.067*	-0.088*	0.005	-0.008	-0.003	0.014	-0.013	0.015	-0.013
Total C (mmol/L)	0.145***	0.009	-0.025	-0.017	-0.311***	-0.147***	0.028	0.063	0.104**	0.070*	0.057	0.015
HDL-c (mmol/L)	-0.222***	-0.057	-0.072*	-0.060*	0.156***	-0.113**	-0.007	-0.010	-0.063	0.005	-0.093**	-0.023
LDL-c (mmol/L)	0.150***	0.039	-0.001	0.005	-0.273***	-0.133**	0.016	0.047	0.093*	0.062	0.056	0.019
Total TG	0.384***	-	0.004	-	-0.543***	-	-0.042	-	0.126***	-	0.141***	-
TG/HDL-c ratio	0.368***	0.016	0.030	0.022*	-0.462***	0.049**	-0.027	0.006	0.121***	0.000	0.140***	0.007
Glucose (mmol/L)	0.106**	0.014	-0.044	-0.049	-0.100**	0.038	-0.027	-0.016	0.035	0.000	0.017	-0.025
HbA1c (%)	0.093**	0.034	-0.020	-0.039	-0.023	0.080	-0.013	-0.021	0.011	-0.022	-0.023	-0.060
Hs-CRP (mg/L) ¹	0.053	-0.038	-0.103*	-0.110*	-0.188***	-0.119*	-0.060	-0.070	-0.047	-0.085	-0.018	-0.065
Creatinine (umol/L)	0.015	-0.038	-0.039	-0.023	-0.067*	0.022	0.018	0.028	0.001	-0.008	-0.001	-0.012
Uric acid (mmol/L) ¹	0.084	-0.134*	-0.150**	-0.163***	-0.230***	0.052	-0.090	-0.085	-0.091	-0.140**	-0.027	-0.079
Ureum (mmol/L) ¹	0.016	0.001	0.130**	0.121*	0.087	0.138*	0.109	0.097	0.143**	0.137*	0.082	0.079

Model S1: Adjusted for age, sex, education and smoking; **Model S2:** Adjusted for age, sex and triglycerides.

Dependent: cardiovascular risk factors, independent: fatty acid status. Associations between fatty acid status and cardiovascular risk factor are reported as standardized β 's. P-value: * = ≤ 0.05 , ** = < 0.01 , *** = < 0.001 . Abbreviations: BMI, body mass index; SBP, systolic blood pressure; DBP, diastolic blood pressure; Total C; total cholesterol; HDL-c, high density lipoprotein cholesterol; LDL-c, Low density lipoprotein cholesterol; Total TG, total triglycerides; TG/HDL-c ratio, total triglycerides/HDL cholesterol; HbA1c, Hemoglobin A1C; Hs-CRP, High sensitive C-reactive protein. ¹ Hs-Crp data was available in n = 412, Uric acid and ureum data were available in n = 348

Table S5: Association between fatty acid markers from plasma phospholipids (PL) and cardiovascular risk factors.

	C14:0		C15:0		C17:0		Trans-C16:1(n-7)		Trans-C18:1(n-7)		CLA	
	Model S1	Model S2	Model S1	Model S2	Model S1	Model S2	Model S1	Model S2	Model S1	Model S2	Model S1	Model S2
Dependent variable												
Weight (kg)	-0.002	-0.015	-0.056	-0.019	-0.150***	-0.102**	-0.011	0.012	-0.056	-0.047	0.012	-0.024
Waist circumference	0.001	-0.023	-0.090**	-0.047	-0.176***	-0.120***	-0.052	-0.027	-0.070*	-0.063*	-0.008	-0.058
BMI (kg/m ²)	-0.018	-0.040	-0.087	-0.017	-0.186***	-0.120**	-0.049	-0.021	-0.099*	-0.087*	-0.025	-0.079*
SBP (mmHg)	-0.026	-0.046	-0.043	-0.017	-0.074*	-0.026	-0.049	-0.041	-0.087*	-0.083*	-0.048	-0.088*
DBP (mmHg)	-0.005	-0.030	-0.067*	-0.038	-0.128***	-0.078*	-0.039	-0.021	-0.063	-0.057	0.012	-0.023
Total C (mmol/L)	0.117**	-	-0.087*	-	-0.253***	-	-0.052	-	0.006	-	0.136***	-
HDL-c (mmol/L)	-0.012	-0.022	0.030	-0.022	-0.063	-0.109***	-0.015	-0.057*	0.022	-0.015	0.018	0.043
LDL-c (mmol/L)	0.120**	0.010	-0.073*	0.005	-0.184***	0.051***	-0.025	0.019	0.011	-0.001	0.100**	-0.028*
Total TG	0.053	-	-0.146***	-	-0.232***	-	-0.114**	-	-0.073*	-	0.120**	-
TG/HDL-c ratio	0.042	0.007	-0.119***	0.010	-0.147***	0.043***	-0.076*	0.025*	-0.061	0.008	0.080*	-0.018
Glucose (mmol/L)	0.029	0.026	-0.060	-0.031	-0.107**	-0.085*	-0.023	0.005	-0.084*	-0.069*	-0.044	-0.065*
HbA1c (%)	-0.019	-0.024	-0.030	-0.019	-0.029	-0.031	-0.018	-0.009	-0.084*	-0.081**	-0.023	-0.037
Hs-CRP (mg/L) ¹	-0.037	-0.056	-0.081	-0.051	-0.222***	-0.200***	-0.110*	-0.102*	-0.172***	-0.168*	0.015	-0.028
Creatinine (umol/L)	-0.041	-0.044	-0.005	0.020	-0.109***	-0.060*	0.002	0.022	-0.055	-0.032	-0.006	-0.022
Uric acid (mmol/L) ¹	-0.087	-0.106*	-0.119*	-0.076	-0.265***	-0.197***	-0.108*	-0.039	-0.197***	-0.128**	-0.047	-0.073
Ureum (mmol/L) ¹	0.012	0.020	0.188**	0.188***	0.091	0.088	0.096	0.104	0.093	0.105	0.028	0.037

Model S1: Adjusted for age, sex, education and smoking; **Model S2:** Adjusted for age, sex, triglycerides and cholesterol.

Dependent: cardiovascular risk factors, independent: fatty acid status. Associations between fatty acid status and cardiovascular risk factor are reported as standardized β 's. P-value: * = ≤ 0.05 , ** = < 0.01 , *** = < 0.001 . Abbreviations: BMI, body mass index; SBP, systolic blood pressure; DBP, diastolic blood pressure; Total C; total cholesterol; HDL-c, high density lipoprotein cholesterol; LDL-c, Low density lipoprotein cholesterol; Total TG, total triglycerides; TG/HDL-c ratio, total triglycerides/HDL cholesterol; HbA1c, Hemoglobin A1C; Hs-CRP, High sensitive C-reactive protein. ¹ Hs-Crp data was available in n = 412, Uric acid and ureum data were available in n = 348

Table S6: Association between dairy fat intake and cardiovascular risk factors

	Dairy fat intake (g/d)					
	Model 1	Model 2	Model 3	Model 4	Model 5	Model 6
Age (years)	0.355***					
Sex	0.083*					
Weight (kg)	-0.003	0.022	0.043	0.037	0.043	0.042
Waist circumference	0.094*	0.007	0.023	0.014	0.023	0.025
BMI (kg/m ²)	0.048	-0.022				
SBP (mmHg)	0.103*	-0.043	-0.048	-0.044	-0.048	-0.046
DBP (mmHg)	0.108**	0.021	0.023	0.012	0.024	0.024
Total C (mmol/L)	0.118**	-0.003	0.006	0.000	0.006	
HDL-c (mmol/L)	0.112**	0.051	0.051	-0.004	0.053	0.036
LDL-c (mmol/L)	0.086*	-0.015	-0.003	0.012	-0.003	-0.011
Total TG	0.009	-0.024	-0.026	-0.009	-0.028	
TG/HDL-c ratio	-0.037	-0.038	-0.039	-0.006	-0.041	-0.014
Glucose (mmol/L)	0.085*	-0.036	-0.024	-0.055	-0.024	-0.018
HbA1c (%)	0.215***	0.020	0.027	0.030	0.027	0.031
Hs-CRP (mg/L) ¹	0.001	0.005	-0.008	0.004	-0.007	-0.012
Creatinine (umol/L)	-0.030	-0.020	-0.026	-0.030	-0.027	-0.023
Uric acid (mmol/L) ¹	-0.172**	-0.122*	-0.107*	-0.106*	-0.101	-0.058
Ureum (mmol/L) ¹	0.188**	0.183**	0.186**	0.184**	0.186**	0.184**

Model 1: Adjusted for energy intake; **Model 2:** Model 1 + age, sex, education and smoking; **Model 3:** Model 1 + age, sex and BMI; **Model 4:** Model 3 + total carbohydrates and total fat; **Model 5:** Model 3 + total fish fat intake; **Model 6:** Model 3 + total triglycerides and total cholesterol.

Dependent: cardiovascular risk factors, independent: dairy fat intake. Associations between dairy fat intake and cardiovascular risk factor are reported as standardized β 's. P-value: * = ≤ 0.05 , ** = < 0.01 , *** = < 0.001 . Abbreviations: BMI, body mass index; SBP, systolic blood pressure; DBP, diastolic blood pressure; Total C; total cholesterol; HDL-c, high density lipoprotein cholesterol; LDL-c, Low density lipoprotein cholesterol; Total TG, total triglycerides; TG/HDL-c ratio, total triglycerides/HDL cholesterol; HbA1c, Hemoglobin A1C; Hs-CRP, High sensitive C-reactive protein. ¹. Hs-Crp data was available in n = 412, Uric acid and ureum data were available in n = 348

Table S7: Association between fish fat intake and cardiovascular risk factors

	Fish fat intake (g/d)					
	Model 1	Model 2	Model 3	Model 4	Model 5	Model 6
Age (years)	0.077*					
Sex	0.004					
Weight (kg)	-0.041	-0.036	0.007	0.003	0.007	0.003
Waist circumference	-0.022	-0.035	0.000	-0.001	0.001	0.007
BMI (kg/m ²)	-0.045	-0.048				
SBP (mmHg)	0.032	0.014	0.013	0.013	0.012	0.022
DBP (mmHg)	0.041	0.020	0.029	0.012	0.029	0.038
Total C (mmol/L)	0.019	-0.015	0.001	-0.011	0.002	
HDL-c (mmol/L)	0.100**	0.080*	0.075*	0.042	0.076*	0.018
LDL-c (mmol/L)	0.018	0.043	0.007	0.009	0.007	-0.001
Total TG	-0.117**	-0.120**	-0.109**	-0.110**	-0.109**	
TG/HDL-c ratio	-0.126***	-0.121***	-0.110***	-0.098**	-0.111***	-0.006
Glucose (mmol/L)	0.000	-0.030	-0.006	-0.014	-0.006	0.016
HbA1c (%)	0.016	-0.026	-0.013	-0.004	-0.012	0.001
Hs-CRP (mg/L) ¹	-0.085	-0.079	-0.047	-0.047	-0.047	-0.033
Creatinine (umol/L)	-0.013	-0.012	-0.015	-0.021	-0.016	-0.002
Uric acid (mmol/L) ¹	0.097	0.102*	0.081	0.079	0.076	0.102**
Ureum (mmol/L) ¹	-0.030	-0.019	-0.012	-0.015	-0.001	-0.011

Model 1: Adjusted for energy intake; **Model 2:** Model 1 + age, sex, education and smoking; **Model 3:** Model 1 + age, sex and BMI; **Model 4:** Model 3 + total carbohydrates and total fat; **Model 5:** Model 3 + total dairy fat intake; **Model 6:** Model 3 + total triglycerides and total cholesterol.

Dependent: cardiovascular risk factors, independent: fish fat intake. Associations between dairy fat intake and cardiovascular risk factor are reported as standardized β 's. P-value: * = ≤ 0.05 , ** = < 0.01 , *** = < 0.001 . Abbreviations: BMI, body mass index; SBP, systolic blood pressure; DBP, diastolic blood pressure; Total C; total cholesterol; HDL-c, high density lipoprotein cholesterol; LDL-c, Low density lipoprotein cholesterol; Total TG, total triglycerides; TG/HDL-c ratio, total triglycerides/HDL cholesterol; HbA1c, Hemoglobin A1C; Hs-CRP, High sensitive C-reactive protein. ¹. Hs-Crp data was available in n = 412, Uric acid and ureum data were available in n = 348

Table S8A: Papers investigating the association between dairy fat biomarkers and cardiovascular risk factors (based on an illustrative search)

Fraction	Plasma/Serum	Paper	Population	n	Outcome	Biomarkers ^{1,2}						
							+/- ₃	C14:0	C15:0	C15:0+ C17:0	C17:0	Trans-C16:1(n-7)
CE+TG+PL+FFA	Plasma	-	-	-	-							
	Serum	Santaren et al. (2014) ¹	Hispanic, African American and Non-Hispanic white adults 40-60 yr	659	Insulin resistance by intravenous glucose-tolerance test (insulin sensitivity index)	r	+		0.14***			-0.12**
						β	+		0.84*			-0.42
					β-cell dysfunction by intravenous glucose-tolerance test (product of the insulin sensitivity index and acute insulin response)	r	+		0.11**			-
						β	+		2.21*			0.10
CE	Plasma	-	-	-	-	-		-	-	-	-	-
	Serum	Smedman et al. (1999) ²	70-year old men	66	Weight	r	+		- 0.36**			
					BMI	r	+		- 0.39***			
					WC	r	+		- 0.28*			
					HC	r	+		- 0.32**			
					HDL triacylglycerol	r	+		- 0.23			
					Insulin sensitivity by clamp (insulin sensitivity index (glucose disposal/actual insulin concentration) *100)	r	+		0.21			
TG	Plasma/Serum	-	-	-	-	-		-	-	-	-	-
PL	Plasma	Mozaffarian et al. (2010) ³	Cardio vascular Health study, adults from 4 US communities	3736	BMI	Δ% ^{a,c}	+					-1.8%*
					WC	Δ%	+					-1.8%***
					LDL	Δ%	0					+2.4%
					HDL	Δ%	+					+1.9%*
					Total Triglycerides	Δ%	+					-19.0%***
					Total:HDL ratio	Δ%	+					-4.7%***
					CRP	Δ%	+					-13.8%*
					Fasting glucose	Δ%	0					-0.96%
					Fasting insulin	Δ%	+					-13.3%***
					Insulin resistance by HOMA-IR	Δ%	+					-16.7%**
		Mozaffarian et al. (2013) ⁴	US adults (45-84 yr) from the Multi-ethnic study of	2617	SBP	Δ% ^{a,c}	+					-2.4%*
					DBP	Δ%	+					-2.9%***
					LDL	Δ%	-					+6.4%***
					HDL	Δ%	0					-1.19
					Total Triglycerides	Δ%	+					-19.1%***

			athero-sclerosis		Total:HDL ratio	$\Delta\%$	0					+1.5%
					Fasting glucose	$\Delta\%$	0					-1.2%
					Fasting insulin	$\Delta\%$	+					-9.1%***
		Nestel et al. (2014) ⁵	Overweight and obese subjects with metabolic syndrome	86	insulin sensitivity index by OGTT (Matsuda index)	β	+				0.30*	0.02
					Insulin resistance (Fasting Insulin) by OGTT	β	+				-2.41*	-0.76
					Insulin resistance (HOMA-IR) by OGTT	β	+				-0.70*	-0.17
		Oliveira et al. (2013) ⁶	US adults (45-84 yr) from the Multi-ethnic study of atherosclerosis	2837	SBP	$\Delta\%^{a,c}$	+	+2.5%*	-4.0%***			-1.18%*
					DBP	$\Delta\%$	+	+1.4%*	-4.1%***			-2.1%**
					LDL	$\Delta\%$	0	+1.9%	-1.11%			+4.4*
					HDL	$\Delta\%$	0	-0.2%	+0.4%			-0.6%
					Total: HDL ratio	$\Delta\%$	0	+5.0%**	-3.4%			+1.0%
					Total Triglycerides	$\Delta\%$	+	+29.5%***	-15.0%***			-17.7%***
					CRP	$\Delta\%$	0	0%	0%			+2.8%
	Serum	Kratz et al. (2014) ⁷	Population without fatty liver disease	32	Fasting glucose	β	+		-1.84*		-0.80*	-1.31*
					AUC glucose by OGTT	β	+		-34.27*		-23.94*	-36.02*
					Incremental AUC glucose by OGTT	β	+		-9.9		-13.26*	-18.68
					Liver fat (liver-spleen ratio)	β	+		2.39		2.13*	3.56*
					Hepatic insulin resistance (basal hepatic glucose * fasting plasma insulin)	β	+		-1.21		-2.68	-5.95*
					Systemic insulin resistance (LL)	β	0		-0.60		5.93	11.2*
					Systemic insulin resistance (HL)	β	0		-0.78		5.72	10.97*
					Pancreatic β -cell function (AIRg)	β	0		2.12		4.35	-1.67
					Pancreatic β -cell function (oral DI)	β	0		2.73		3.82	4.34
					Pancreatic β -cell function (IV DI)	β	0		1.68		6.24	1.24
		Rosell et al. (2004) ⁸	Healthy men (63-year old)	299	Abdominal obesity	r	+	0.09	-0.22***		-0.30***	
		Smedman et al. (1999) ²	Men (70-years old)	66	Weight	r	0		-0.21			
					HC	r	+		-0.25*			
					Apo-B	r	0		-0.27			
		Warensjö et al. (2010) ⁹	Swedish general population	1000	SBP	r	+		-0.05	-0.10***	-0.11***	
					DBP	r	+		-0.06	-0.13**	-0.16**	
					BMI	r	+		-0.08*	-0.14**	-0.14**	
					Total cholesterol	r	+		-0.10***	-0.04	0.008	

					Total Triglycerides	r	+		-0.18**	-0.23**	-0.22*	
					Fasting glucose	r	+		-0.05	-0.13**	-0.16*	
FFA	Plasma/Serum	Kratz et al. (2014) ⁷	Population without fatty liver disease	32	Fasting glucose	β	+		-1.09*		-0.54	-1.23
					AUC glucose by OGTT	β	+		-32.43*		-18.07*	-35.25
					Incremental AUC glucose by OGTT	β	+		-18.00*		-10.96*	-18.95
					Liver fat (liver-spleen ratio)	β	+		2.55*		1.80*	3.20
					Hepatic insulin resistance (basal hepatic glucose * fasting plasma insulin)	β	0		-2.21		-0.74	-4.79
					Systemic insulin resistance (LL)	β	0		7.24		1.15	7.95
					Systemic insulin resistance (HL)	β	0		6.89		1.10	8.14
					Pancreatic β-cell function (AIRg)	β	0		-2.32		2.29	-3.49
					Pancreatic β-cell function (oral DI)	β	0		3.75		1.30	-1.82
					Pancreatic β-cell function (IV DI)	β	0		2.15		4.05	4.82
Erythrocytes	-	-	-	-	-	-	-	-	-	-	-	-
AT	-	Castro-Webb et al. (2012) ¹⁰	Costa-Rican adults	1744 (232 cases)	Type 2 diabetes	PR ^b	0			1.33 (0.88-2.11)		1.05 (0.66-1.66)
		Iggman et al. (2010) ¹¹	Elderly men	795	Insulin sensitivity by clamp (glucose infusion rate)	r	+	0.32***	0.07		0.21***	
						β	+	0.35**	0.04		3.5*	
					Insulin sensitivity (HOMA-IR)	r	+	-0.27***	-0.04		-0.21***	
						β	+	-0.10*	0.10		-1.4***	
		Rosell et al. (2004) ¹²	Healthy men (63-year old)	297	Abdominal obesity	r	+	-0.30***	-0.20***		-0.31***	
		Smit et al. (2010) ¹³	Costa-Rican Adults	1785	BMI	β	-					0.37***
					WC	β	-					0.95***
					SS	β	+					-2.27***
					TS	β	+					-1.38***
					SUS	β	+					-1.67***

Significance level: * = p < 0.05, ** = p < 0.01, *** = p < 0.001.

^a. Relationships of the dairy fatty acid with cardiovascular risk factors were evaluated using linear regression; ^b. Highest versus lowest quintile; ^c. Δ% is calculated by the authors of this manuscript, using data from the study.

¹. r = correlation coefficient, Δ% = difference between low and high fatty acid biomarker, β = beta-coefficient, PR = prevalence ratio

². Adjustments: Santaren (2014) = demographics, lifestyle, dietary variables; Mozaffarian (2010) = age, sex, diabetes, smoking, coronary heart disease, physical activity, dietary intake (BMI, waist circumference); Mozaffarian (2013) = age, sex, race, education, smoking, alcohol intake, physical activity, prevalent diabetes, waist circumference; Nestel (2014) = age, systolic blood pressure,

waist:hip ratio; Oliveira (2013)= demographics, lifestyle, prevalent diabetes, medication; Kratz (2014)= age, sex, BMI; Castro-Webb (2012)= demographics, lifestyle, several fatty acids; Iggman (2010)= BMI, smoking, alcohol intake, physical activity; Smit (2010)= age, sex, residence, all trans-fatty acids, smoking, physical activity, income, diabetes, hypertension, energy intake and adipose tissue alpha-linolenic acid

³. + = An increase of the biomarker reduces the cardiovascular risk factor. - = An increase in the biomarker increases the cardiovascular risk factor. 0 = No significant effect

Abbreviations: AIRg, acute insulin response to glucose; AT, Adipose tissue; AUC, area under the curve for glucose; BMI, Body mass index; C14:0, Myristic acid; C15:0, Pentadecanoic acid; C17:0, Heptadecanoic acid; CE, Cholesterol esters; CRP, C-reactive protein; DBP, diastolic blood pressure; FFA, Free fatty acids; HC, Hip circumference; HDL, High density lipoprotein; HL, High level insulin infusion; HOMA-IR, homeostatic model assessment-estimated insulin resistance; IV DI, intravenous disposition index; LDL, Low density lipoprotein; LL, low-level insulin infusion; OGTT, oral glucose tolerance test; oral DI, oral disposition index; PL, Phospholipids; SBP, systolic blood pressure; SS, Subscapular Skinfold; SUS, Suprailiac skinfold; TG, Triglycerides; Trans-C16:1(n-7), Trans-Palmitoleic acid; TS, Triceps skinfold; WC, waist circumference.

Table S8B: Papers investigating the association of dairy fatty acid biomarkers with the development of cardiovascular diseases (based on an illustrative search)

Paper	Population	n	Outcome	Blood Sample	Lipid Fraction		+/- ³	Biomarkers ^{1,2} (Risk (95%CI))			
								C14:0	C15:0	C17:0	Trans-C16:1(n-7)
			Coronary Heart Disease								
Iggman et al. (2016) ¹⁴	Uppsala Longitudinal Study of Adult Men (ULSAM)	535 (251 cases)	Cardiovascular diseases	Adipose Tissue		HR ^a	0	0.95 (0.79 – 1.15)	1.03 (0.86-1.23)	0.98 (0.82-1.16)	
Khaw et al. (2012) ¹⁵	Men and women from the EPIC cohort study	2424 cases, 4930 controls	Incidence of coronary heart disease	Plasma	PL	OR ^a	0	0.97 (0.87-1.08)	0.98 (0.89-1.09)	0.91 (0.82-1.01)	
Matthan et al. (2014) ¹⁶	Post-menopausal women	2448 (1224 cases, 1224 controls)	Coronary heart disease	Plasma	PL	OR ^a	0	1.05 (0.70-1.56)	1.02 (0.61-1.72)		
Oliveira et al. (2013) ⁶	US adults (45-84yr) from the Multi-ethnic study of atherosclerosis	2837 (189 cases)	Incidence of cardiovascular disease	Plasma	PL	HR ^b	0	1.06 (0.65-1.74)	0.55 (0.32-0.95)		0.99 (0.58-1.69)
						HR ^a	+	1.02 (0.87-1.20)	0.81* (0.68-0.98)		0.97 (0.82-1.13)
		2837 (146 cases)	Incidence of coronary heart disease	Plasma	PL	HR ^b	+	1.13 (0.63-2.03)	0.41* (0.22-0.78)		0.82 (0.45-1.49)
						HR ^a	+	1.01 (0.84-1.21)	0.74* (0.60-0.92)		0.91 (0.75-1.10)
Sun et al. (2007) ¹⁷	Female nurses, 30-55 years old	166 cases, 327 controls	Ischemic heart disease	Plasma	Total	RR ^c	-		2.36 * (1.16-4.78)	1.25 (0.68-2.30)	0.79 (0.44-1.42)
				Erythrocytes		RR ^c	0		0.93 (0.42-2.05)	0.72 (0.38-1.38)	0.98 (0.53-1.83)
Warensjö et al. (2010) ⁹	Swedish general population	444 cases (307 men), 556 controls (308 men)	First myocardial infarction	Plasma	PL	OR ^a (men)	0		0.92 (0.78-1.10)	0.92 (0.78-1.10)	
				Plasma	PL	OR ^a (women)	+		0.78* (0.61-0.99)	0.74* (0.58-0.95)	
			Stroke								
Warensjö et al. (2009) ¹⁸	Northern Swedish general population	108 stroke cases, 216 controls	First stroke	Plasma	PL	OR ^a	+		0.81* (0.60-1.10)	0.71* (0.52-0.79)	
Yaemsiri et al. (2013) ¹⁹	Womens Health Initiative	964 cases, 964 controls	Incident ischemic stroke	Serum	PL	OR ^{b,d}	0	1.09 (0.86-1.37)	1.07 (0.87-1.13)	1.01 (0.85-1.21)	0.94 (0.76-1.16)

	Observational Study (WHI-OS)										
Yakoob et al. (2014) ²⁰	Women in the Nurses' Health study (NHS) and men in the Health professionals study (HPFS)	472 cases NHS, 122 cases HPFS, 594 controls	Incident stroke	Plasma	Total	HR ^e	0	1.05 (0.62-1.78)	0.85 (0.54-1.33)	0.99 (0.67-1.49)	0.89 (0.55-1.45)
				Erythrocytes		HR ^e	0	1.31 (0.73-2.36)	0.88 (0.53-1.45)	0.87 (0.42-1.76)	1.07 (0.61-1.88)
Yamagishi et al. (2013) ²¹	Men and Women from the Minneapolis Field Center of the Atherosclerosis Risk in Communities Study (ARIC)	3870 (168 cases)	Incident Ischemic stroke	Plasma	CE	HR ^e	0	1.43 (0.92-2.22)	0.83 (0.55-1.25)		
					PL	HR ^e	0	1.29 (0.83-2.01)	0.89 (0.60-1.33)		
			Heart Failure								
Matsumoto et al. (2013) ²²	Physicians Health Study (PHS)	788 cases, 788 controls	Incident heart failure	Plasma	PL	OR ^e	0	1.03 (0.65-1.64)	0.73 (0.44-1.12)		
						OR ^a	0	1.02 (0.86-1.20)	0.91 (0.76-1.07)		
Tokede et al. (2013) ²³	Physicians Health Study (PHS)	788 cases, 788 controls	Incident heart failure	Plasma	PL	OR ^b	0				0.74 (0.49-1.10)
						OR ^a	0				0.91 (0.80-1.04)
Yamagishi et al. (2008) ²⁴	Men and Women from the Minneapolis Field Center of the Atherosclerosis Risk in Communities Study (ARIC)	3592 (197 cases)	Incident heart failure	Plasma	CE	HR ^e	0	1.70** (1.06-2.71)	0.80 (0.50-1.28)	0.81 (0.52-1.26)	
				Plasma	PL	HR ^e	+	1.29 (0.80-2.08)	0.62* (0.38-1.02)	0.55** (0.35-0.85)	

Significance level: * = p < 0.05, ** = p < 0.01, *** = p < 0.001. One asterisk was ascribed when the risk was significant, but the level unknown.

^a. Evaluated as continuous variables (per 1 SD unit difference)^b. Highest versus lowest quintile ^c. Highest versus lowest tertile ^d. CI is 99.9% instead of 95%, ^e Highest versus lowest quartile.

¹. RR= Relative risk, HR= hazard ratio, OR= Odds ratio

². Adjustments: Iggman (2016)= age, sex and cardiovascular risk factors; Kwah (2012)= age, sex, polyunsaturated fatty acids, BMI, smoking, alcohol intake, physical activity, social class, education, blood pressure; Matthan (2014)= demographics, BMI, systolic blood pressure, smoking, education, medication and hormone use, family history of cardiovascular disease/stroke/myocardial infarction/type 2 diabetes, physical activity, dietary intake; de Oliveira (2013)= demographics, lifestyle, dietary variables; Sun (2007)= age, smoking, fasting status, BMI, postmenopausal status and hormone use, physical activity, alcohol intake, aspirin intake, history of hypertension, hypercholesterolemia, diabetes, linoleic acid, total trans fatty acids; Warensjo (2009) = BMI, serum-cholesterol, tobacco use, systolic blood pressure, diastolic blood pressure; Yaemsiri (2013)= BMI, smoking, diabetes, aspirin use, systolic blood pressure, antihypertensive medication, total cholesterol, HDL cholesterol and triglycerides; Yakoob (2014)= age, race, month of blood collection, smoking status, physical activity, alcohol, family history of diabetes, myocardial infarction, menopausal status, postmenopausal hormone use, food intake, energy intake, plasma/erythrocytes trans 18:1, trans 18:1, 16:0, 18:0; Yamagishi (2008 & 2013)= Age and sex; Matsumoto (2013) and Tokede (2013)= age, race, year of birth, blood sample collection, BMI, smoking, exercise, alcohol intake, history of diabetes and atrial fibrillation and saturated fatty acids.

³. + = An increase of the biomarker decreases the risk of diabetes type 2 and/or cardiovascular diseases. - = An increase in the biomarker increases the risk of diabetes type 2 and/or cardiovascular diseases. 0 = No significant effect

Abbreviations: AT, Adipose tissue; C14:0, Myristic acid; C15:0, Pentadecanoic acid; C17:0, Heptadecanoic acid; CE, Cholesterol esters; FFA, Free fatty acids; PL, Phospholipids; TG, Triglycerides; Trans-C16:1(n-7), Trans-Palmitoleic acid.

Table S8C: Papers investigating the association of dairy fatty acid biomarkers with the development of diabetes type 2 (based on an illustrative search)

Paper	Population	n	Blood Sample	Lipid Fraction		+/- 3	Biomarkers ^{1,2} (Risk (95%CI))			
							C14:0	C15:0	C17:0	Trans-C16:1(n-7)
Forouhi et al. (2014) ²⁵	Adults in the EPIC cohort study	27296 (12132 cases, 15164 controls)	Plasma	PL	HR ^a	+	1.15* (1.09-1.22)	0.79* (0.73-0.85)	0.67* (0.63-0.71)	
Hodge et al. (2007) ²⁶	Adults in the Melbourne collaborative cohort study	364 cases, 3391 controls	Plasma	PL	OR ^b	+		0.26*** (0.17-0.40)		
Krachler et al. (2008) ²⁷	Adults in the VIP study	159 cases, 291 controls	Erythrocytes		OR ^a	+	1.25* (1.01-1.53)	0.65** (0.50-0.85)	0.47*** (0.35-0.65)	
Mozaffarian et al. (2010) ³	Cardio-vascular Health study, adults	2985 (304 cases)	Plasma	PL	HR ^b	+				0.38*** (0.24-0.62)
					HR ^c	0		0.99 (0.89-0.10)	0.98 (0.85-1.13)	
Mozaffarian et al. (2013) ⁴	US adults (45-84yr) from the Multi-ethnic study of atherosclerosis	2281 (205 cases)	Plasma	PL	HR ^b	+	0.98 (0.63-1.53)	0.70 (0.42-1.15)		0.52* (0.32-0.85)
Santaren et al. (2014) ¹	Hispanic, African American and Non-Hispanic white adults 40-60 yr	659	Serum	Total	OR ^a	+		0.73* (0.56-0.95)		
Yakoob et al. (2016) ²⁸	Women in the Nurses' Health study (NHS) and men in the Health professionals study (HPFS)	3333 (184 cases NHS study, 93 cases HPFS study)	Plasma	Total	HR ^d	+	1.03 (0.65-1.64)	0.56* (0.37-0.86)	0.57* (0.39-0.83)	0.48*** (0.33-0.70)
		3289 (179 cases NHS study, 97 cases in HPFS study)	Erythrocytes		HR ^d	+	1.56 (0.98-2.49)	0.83 (0.55-1.25)	0.54*** (0.40-0.73)	0.78* (0.51-1.18)

Significance level: * = p < 0.05, ** = p < 0.01, *** = p < 0.001. One asterisk was ascribed when the risk was significant, but the level unknown.

^a. Evaluated as continuous variables (per 1 SD unit difference), ^b. Highest versus lowest quintile, ^c. HR's evaluated per 0.05% points of total fatty acids ^d. Highest versus lowest quartile

¹ RR= Relative risk, HR= hazard ratio, OR= Odds ratio

². Adjustments: Forouhi (2014), Santaren (2014), Mozaffarian (2010, 2013)= demographics, lifestyle, dietary variables; Hodge (2007)= demographic, diabetes history, physical activity, alcohol intake; Krachler (2008)= alcohol intake; Yakoob (2016)= age, race, month of blood collection, smoking status, physical activity, alcohol, family history of diabetes, myocardial infarction, menopausal status, postmenopausal hormone use, food intake, energy intake, plasma/erythrocytes trans 18:1, trans 18:1, 16:0, 18:0;

³. + = An increase of the biomarker decreases the risk of diabetes type 2 and/or cardiovascular diseases. - = An increase in the biomarker increases the risk of diabetes type 2 and/or cardiovascular diseases. 0 = No significant effect

Abbreviations: AT, Adipose tissue; C14:0, Myristic acid; C15:0, Pentadecanoic acid; C17:0, Heptadecanoic acid; CE, Cholesterol esters; FFA, Free fatty acids; PL, Phospholipids; TG, Triglycerides; Trans-C16:1(n-7), Trans-Palmitoleic acid.

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