

Genetic variation in *APOJ*, *LPL*, and *TNFRSF10B* affects plasma fatty acid distribution in Alaskan Eskimos<sup>1-3</sup>

1. 1994-1998, CVD 30-40%  
Al-N (25-54)  
U.S. (9). S. 1  
Al-N US  
(9). G.

SNP (15),  $\sigma^2_{\text{SOLAR}} = 4.0$

FA (16)

$$P = G\sqrt{h_1^2}\sqrt{h_2^2} + E\left[\sqrt{(1-h_1^2)}\sqrt{(1-h_2^2)}\right]$$

(16)

T (17)

(17)

0.

### Measured genotype analysis

SNP (18)

LD (19)

SOLAR (19)



1,115 SNP  
SNP LD (>90%) SNP . T

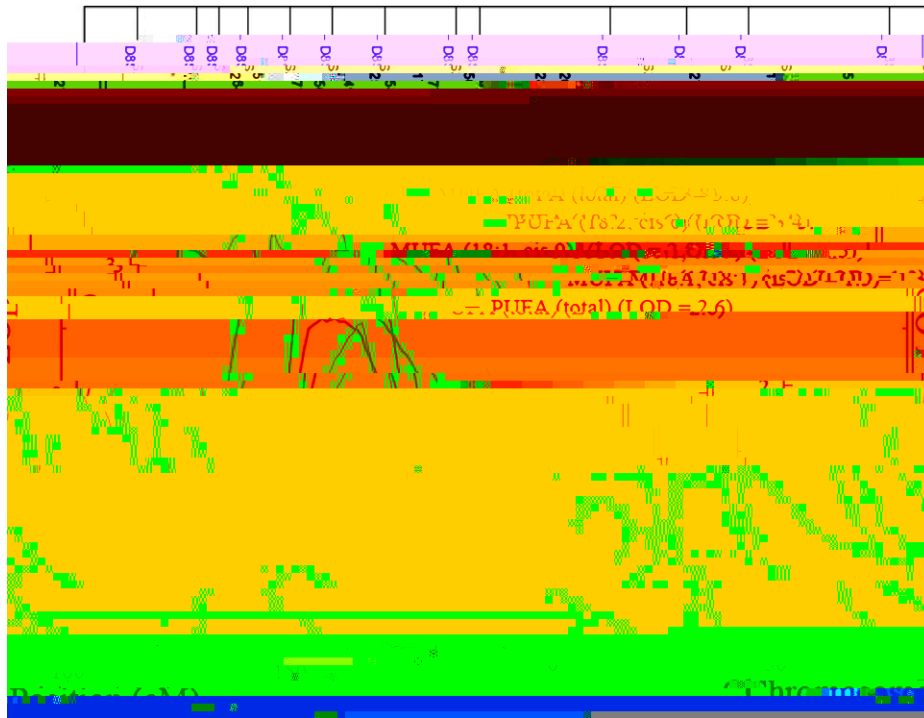


FIGURE 2.  $E_{\text{LIPID}}^{\text{V}}$  (LOD) scores for lipid traits across chromosomes 1-22. Significant associations are highlighted with red and orange bars. Key traits include PUFA (18:2 cis) (LOD=14.1), MUFA (18:1 cis) (LOD=11.7), PUFA (18:3 cis) (LOD=11.2), and PUFA (total) (LOD=2.6).

### DISCUSSION

In this study, we identified a QTL for PUFA on chromosome 8 (8:12-21) that is associated with PUFA. This QTL is associated with PUFA (total) (LOD=2.6) and PUFA (18:2 cis) (LOD=14.1). The QTL is associated with PUFA (total) (LOD=2.6) and PUFA (18:2 cis) (LOD=14.1).

TABLE 3

Summary statistics for lipid traits across chromosomes 1-22 (n = 761)

Trait	Trait	$\beta$ ± SE	$\beta$ ± SE
Metabolic	BMI	0.552 ± 0.17	0.257 ± 0.04
	Weight	0.433 ± 0.17	0.241 ± 0.04
	Percentage	0.601 ± 0.17	0.260 ± 0.04
	Triglyceride	0.690 ± 0.26	0.244 ± 0.05
	HDL cholesterol	-0.36 ± 0.15	-0.350 ± 0.03
Lipid	Triglyceride	0.732 ± 0.13	0.563 ± 0.03
	Glucose	0.565 ± 0.23	0.193 ± 0.04
	Triglyceride	0.329 ± 0.13	0.362 ± 0.03
Statistical	Percentage	-0.357 ± 0.14	-0.165 ± 0.04
	Triglyceride	-0.39 ± 0.19	-0.123 ± 0.04
	HDL cholesterol	0.41 ± 0.17	0.124 ± 0.04
	Triglyceride	-0.543 ± 0.25	-0.156 ± 0.05
	Triglyceride	-0.57 ± 0.12	-0.316 ± 0.04
MUFA, 18:1 -9	Triglyceride	-0.387 ± 0.0837	0.0837

TABLE 4

β; l . . . . . 1 μ . . . . . 0.5 t . . . . . μ . . . . . 8 0.5 l μ . . . . . 0.5 t b . . . . . 0.5 . . . . . t - l t . . . . . t . . . . .

( = 761)

G	SNP	TFA	MUFA	PUFA	SFA	18:1 7	18:1 -9	18:2 -6	BMI	% F t	W t	T . l . 0.5
	35361594	0.226	0.267	0.371	0.767	0.304	0.413	0.206	0.025	0.009	0.043	0.545
	10503814	0.173	0.028	0.017	0.619	0.727	0.381	0.037	0.414	0.165	0.799	0.742
	11136000	0.604	0.364	0.802	0.382	0.509	0.611	0.368	0.002	0.005	0.003	0.04
	1982229	0.992	0.409	0.742	0.916	0.097	0.041	0.388	0.252	0.555	0.126	0.135
	538181	0.033	0.076	0.274	0.584	0.197	0.841	0.279	0.161	0.065	0.600	0.784
	569205	0.032	0.110	0.263	0.505	0.512	0.910	0.385	0.095	0.022	0.390	0.859
	7812347	0.874	0.016	0.722	0.657	0.080	0.024	0.883	0.128	0.032	0.057	0.502
	9331891	0.008	0.188	0.057	0.331	0.295	0.218	0.230	0.780	0.688	0.986	0.073
	1059611	0.172	0.086	0.828	0.975	0.376	0.994	0.041	0.718	0.517	0.253	6.4 × 10 <sup>-6</sup>
	1121923	0.380	0.199	0.467	0.565	0.863	0.615	0.177	0.024	0.011	0.034	0.636
	13702	0.745	0.023,2(0.331)-2 0.5			0.565	0.863T 0.3272					× 10

T t b t . . . . . l . . . . . t . . . . . t . . . . . l . . . . . t . . . . . H t . . . . . ll , Al . . . . . E μ . . . . . v ; . . . . . μ . . . . .

0.5 t . . . . . t . . . . . t . . . . . t . . . . . fl . . . . . ll . . . . . μ . . . . . -3 FA b . . . . . t . . . . . t . . . . . 0.5 t . . . . . l . . . . . It . . . . .

l t , FA t . . . . . 0.5 t . . . . . t . . . . . t . . . . . b t . . . . . v . . . . . l μ FA . . . . . t . . . . . t . . . . . l . . . . . CVD μ . . . . . t . . . . . t . . . . .

0.5 SNP . . . . . t . . . . . l . . . . . 0.5 t . . . . . t . . . . . t . . . . . v ; . . . . . μ . . . . . t . . . . . l t . . . . . (8). H v ; . . . . . t . . . . .

t . . . . . l . . . . . t . . . . . SNP . . . . . l μ FA . . . . . t . . . . . v ; . . . . . μ . . . . . CVD μ . . . . . t . . . . . t . . . . . A . . . . .

PI μ FA 0.5 t b . . . . . 0.5 t . . . . . t . . . . . l μ SFA t . . . . . t . . . . . t . . . . . l . . . . . 0.5 . . . . . t . . . . . t . . . . . t . . . . .

MUFA . . . . . PUFA t . . . . . t . . . . . t . . . . . t . . . . . μ . . . . . 0.5 . . . . . t . . . . . l . . . . . t . . . . . v ; . . . . . μ . . . . . t . . . . . fl . . . . . t . . . . .

t . . . . . l μ SFA 0.5 l . . . . . t . . . . . t . . . . . PUFA . . . . . t . . . . . CVD . . . . . l (8).

TABLE 5

B: QTLs for BMI (BQTN) in the 8

T1	G	SNP	M11/	M11	P11	L11	SNP
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T11 569205 A/0.484 0.74 0.68 F1, 5' UTR

B: fl... FA... S... FA... fl... FA... Alt... I... FA... (5, 24).  
H... 0.35... 0.55... T... PUFA... T... 1(6) ... 0.28 ... 0.38...  
G: ... FA... CVD... T2DM, ... QTL... MUFA... 8, ... -LOD... ≈20... M... T... QTL... PUFA, ... M... β3-... ( ), L... ( ), V... J... ( ), ... 10B ( ). ... G-... (ADRB3) ... (25). A... T... 26A, ... (26, 27). Alt... SNP... FA... BMI, ... T... SNP, 1... 5'... Alt... BMI, ... (28). ... FA (29).

A: 291S ... (30). F... M... C... (27) ... SNP... BMI... I... 285... PUFA 18:2 -6. T... II... (31). Alt... SNP (1... )... LD... A... SNP, 328, ... S 447. I... SNP... (G) ... N... HDL... (36). M... (32). -LOD... HDL... P... Al... (33-35). I... HDL... (36, 37). I... SNP 9331891, ... FA, ... FA... G... I... (36). Al... -LOD... ( )/4... (TNF) ... 5 ( )/4... 2... TNFSRF10B, ... ll1... FADD (38). F... FA...





29. M... MG, E... J, C... JC, ... I. A... ... L...  
... (LPL)... ... 8 22. C... C II  
G... 1993;63:45-6.
30. H... Y, L... W, H... R, Z... X. A... V;  
... L... A 291S;  
... J. L... R... 2006;47:1908-14.
31. S... GS, T... I, S... I, G... I. S... L... L... I-  
... L... : H. GE...  
... Au J E... 2008;168:1233-46.
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S447X SNP...  
2001;60:293-300.
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... E... R... C... 2010;17:R1-17.
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... CLU... PICALM... Al-  
... N... G... 2009;41:1088-93.
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... CR1... Al...  
... N... G... 2009;41:1094-9.
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... I V... 2008;22:537-42.
37. N... CS, B... CH, S... DK, A... CE, U... I FA,  
K... MI. A... J... HDL...  
... A... H... B... 1999;71:197-218.
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