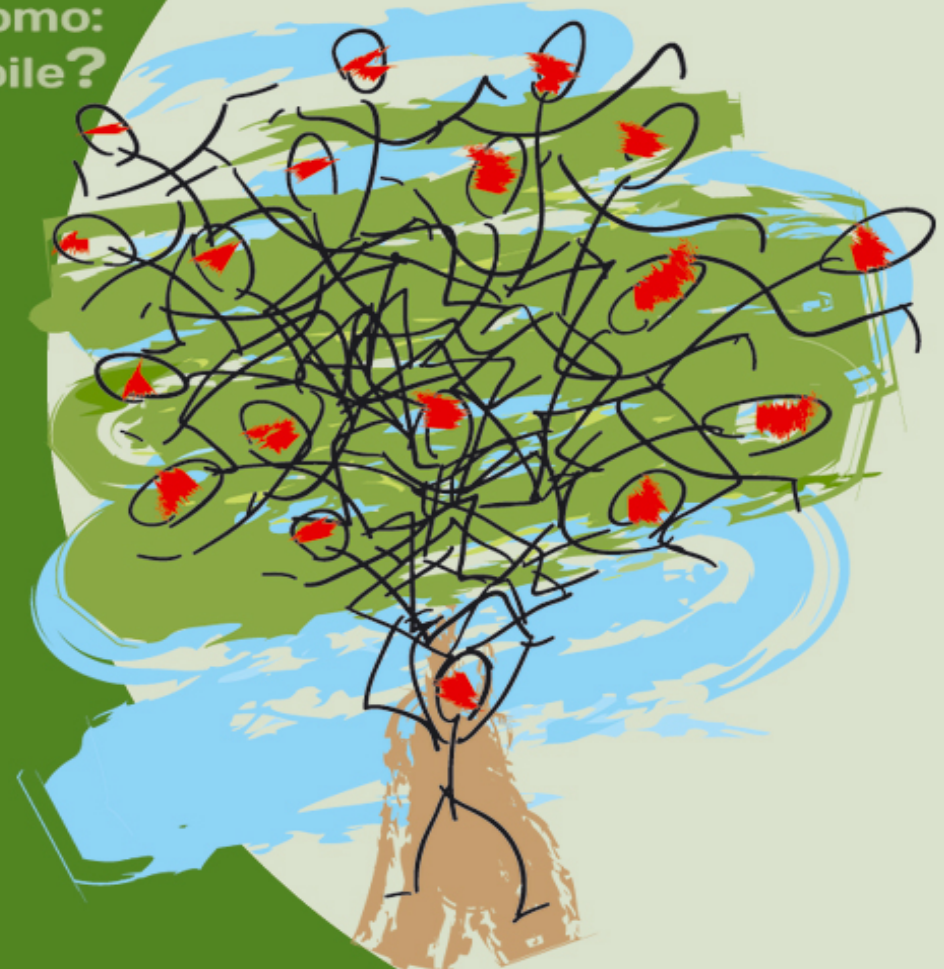


# GLOBALIZZAZIONE E MALATTIE CRONICHE

dall'evidenza scientifica alla cultura dell'uomo:  
un passo impossibile?

## Lo stile di vita per le dislipidemie

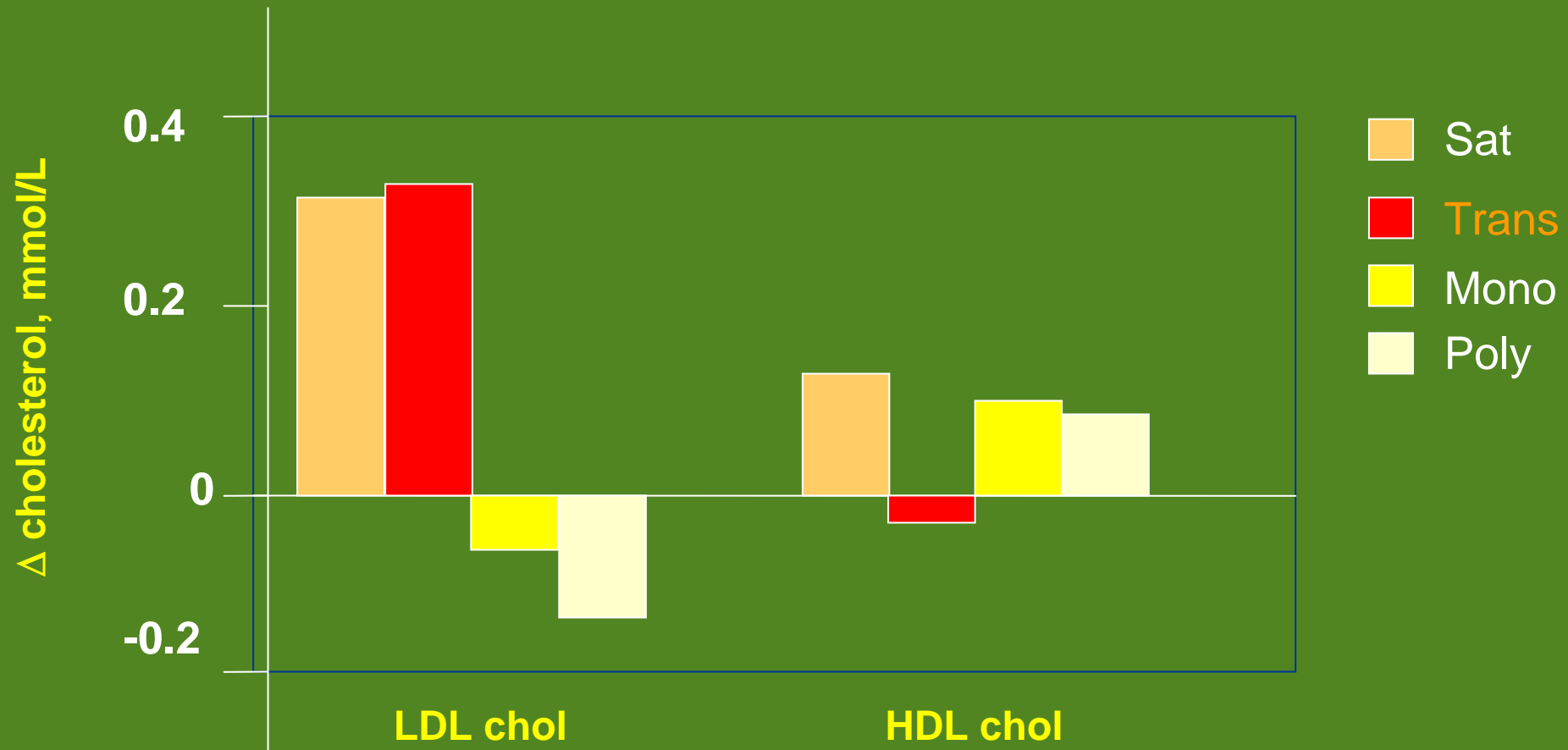
**Andrea Poli**  
*Nutrition Foundation of Italy*



SCUOLA NAZIONALE DI MEDICINA DEGLI STILI DI VITA  
FIMMG - METIS  
AULA POCCHIARI - Istituto Superiore di Sanità - 09 Aprile 2010

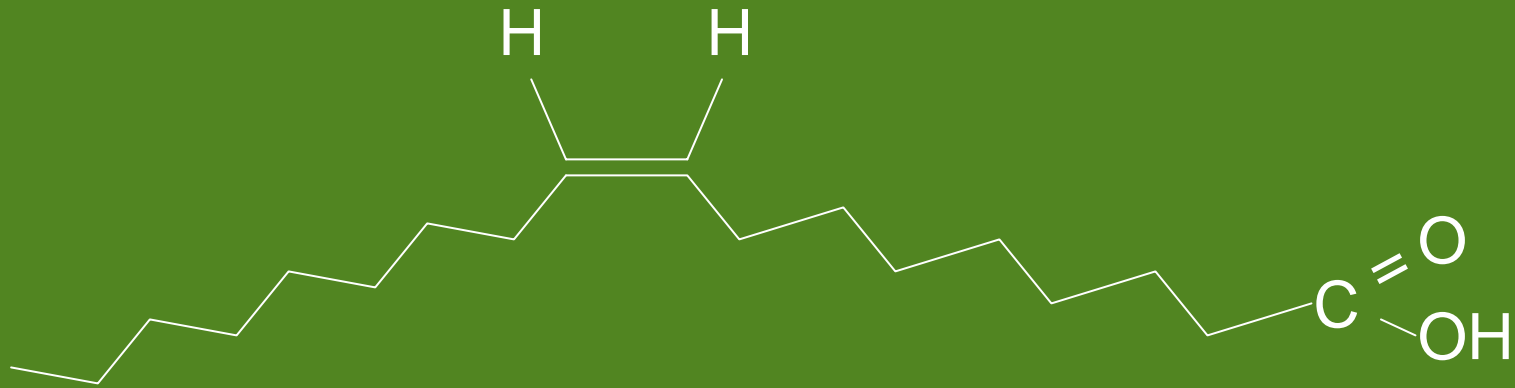


## Effects of SAT, *trans* MONO, *cis* MONO, and *cis* POLY Fatty Acids on LDL and HDL Cholesterol

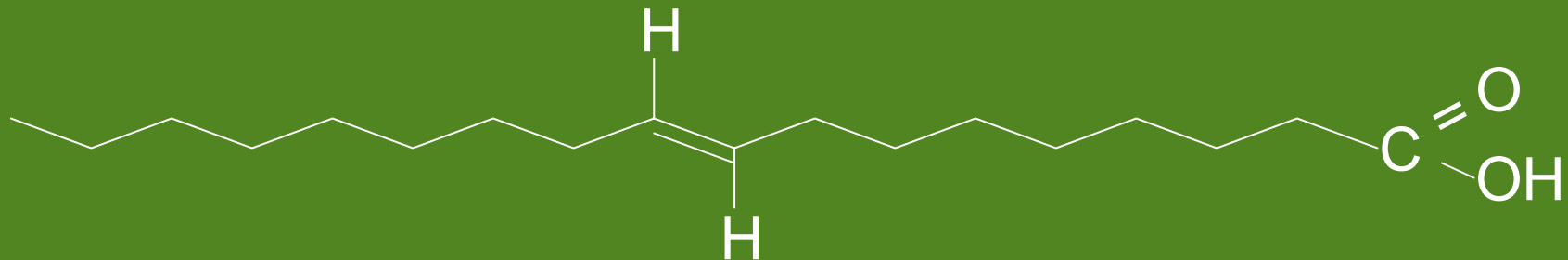


Values obtained by meta-analysis of 32 controlled dietary trials in humans

# Structure of *cis* and *trans* Fatty Acids

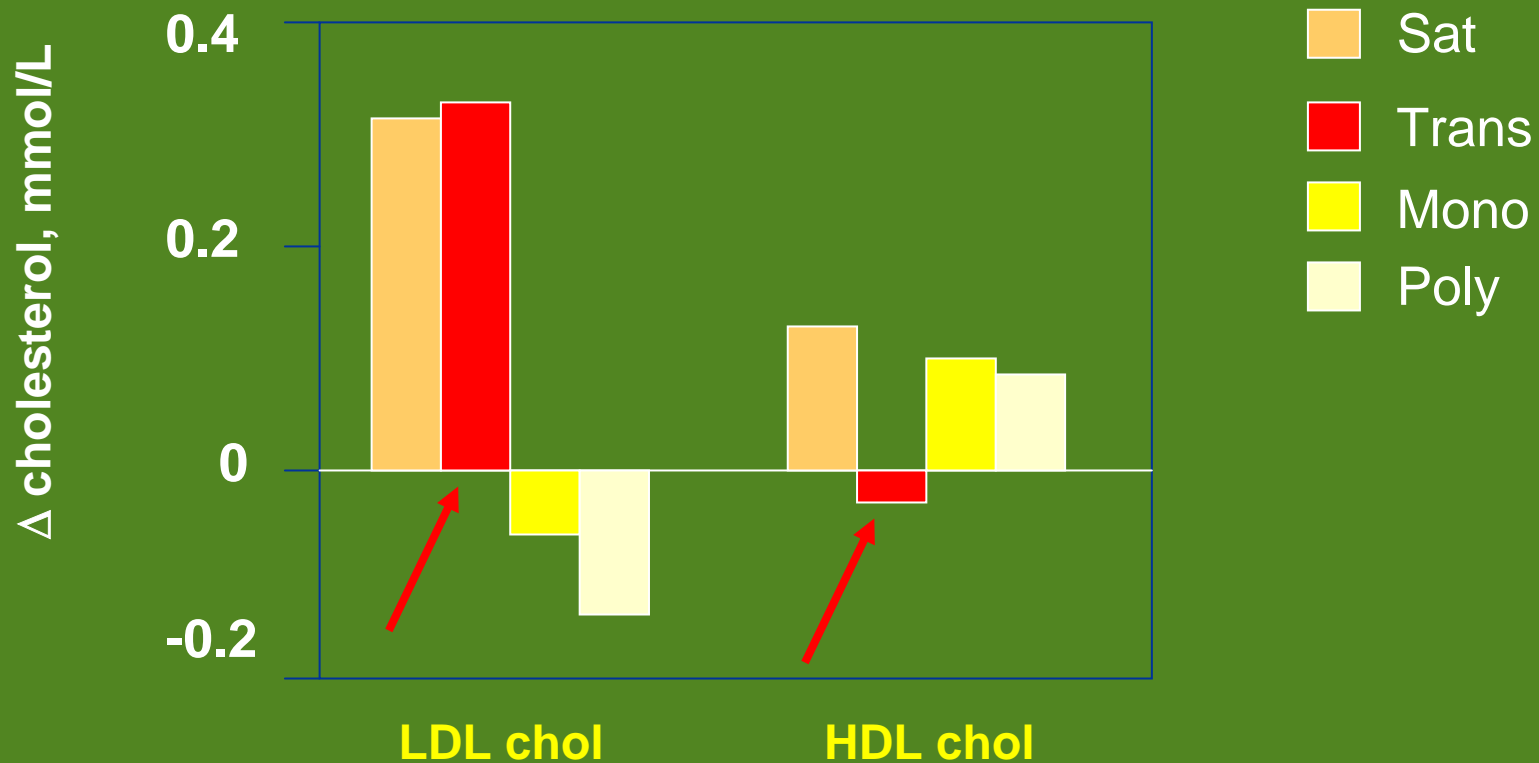


*cis* double bond: oleic acid



*trans* double bond: elaidic acid

# Effects of SAT, *trans* MONO, *cis* MONO, and *cis* POLY Fatty Acids on LDL and HDL Cholesterol



Values obtained by meta-analysis of 32 controlled dietary trials in humans

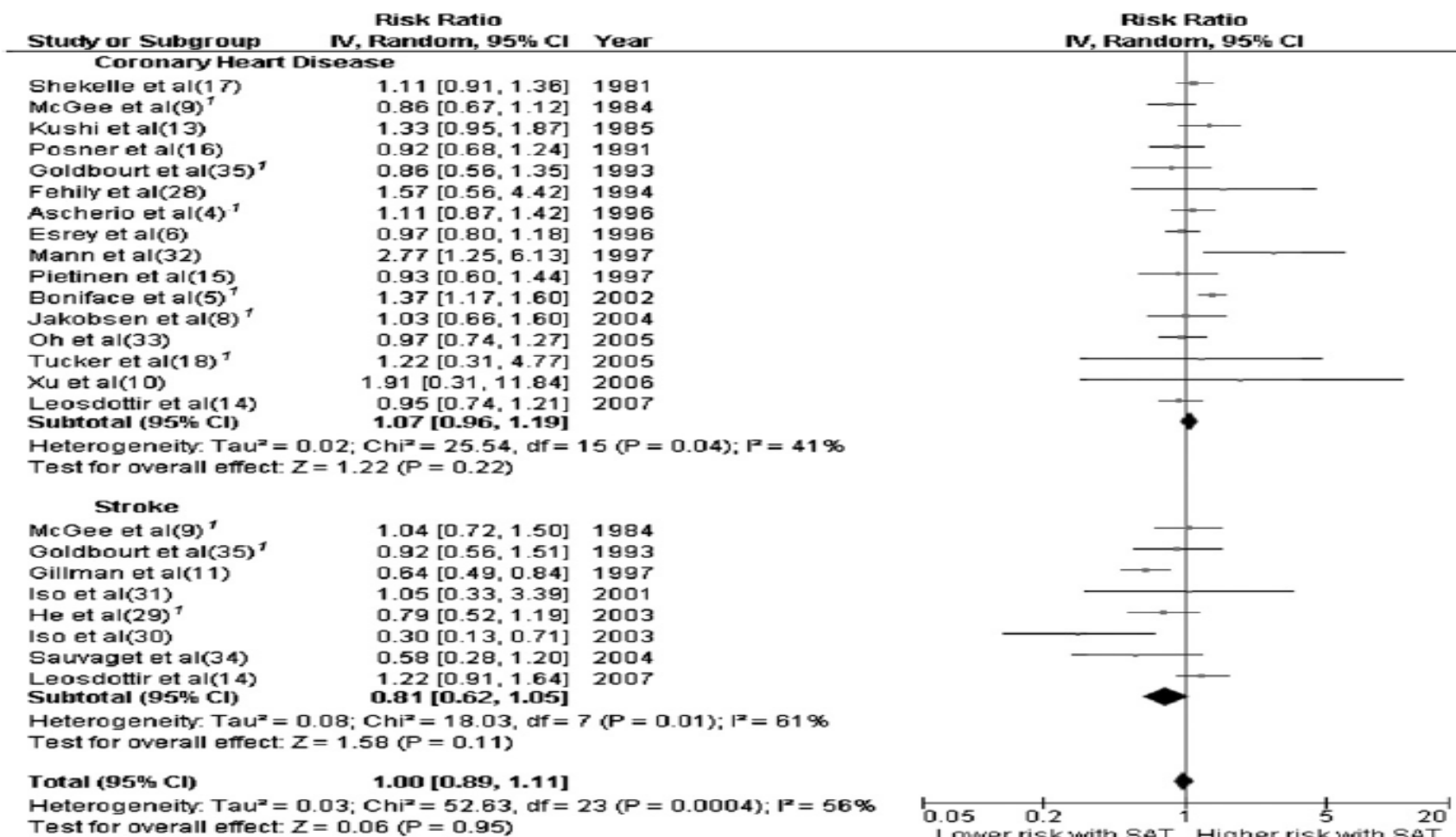
Values obtained by meta-analysis of 32 controlled dietary trials in humans

## Biomarkers of inflammation and endothelial dysfunction and trans fatty acid intake in the Nurses' Health Study (1986-1990)

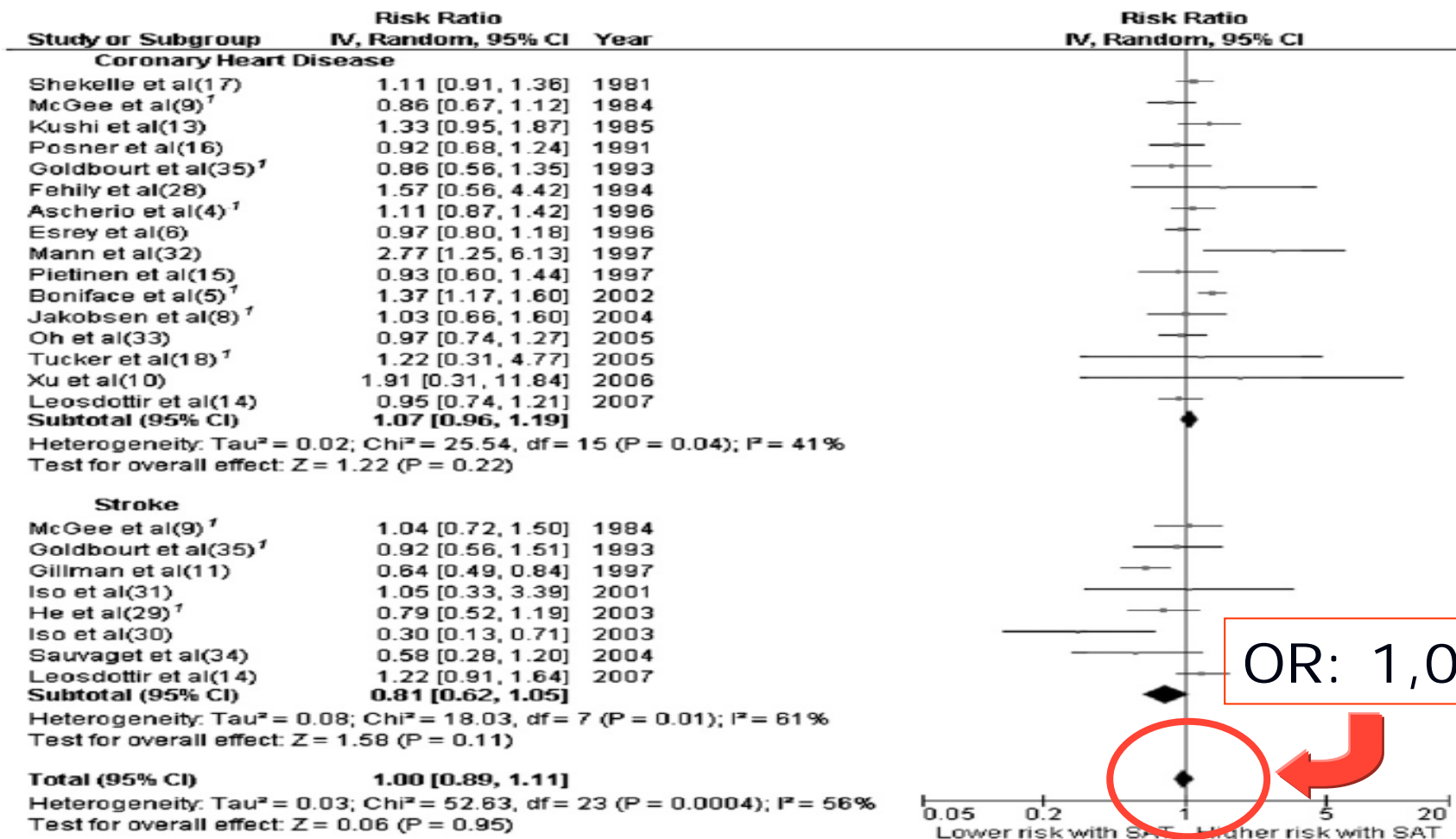
Quintile	<i>n</i>	<i>CRP mg/L</i>	<i>IL-6 ng/L</i>	<i>E-selectin ng/L</i>
Trans fatty acids (range; g/d)				
Q1 (0.61-1.87)	147	1.1 (0.9, 1.3)	1.8 (1.6, 2.0)	41.8 (39.0, 44.9)
Q2 (1.88-2.26)	145	1.3 (1.1, 1.6)	1.7 (1.5, 2.0)	41.9 (39.0, 45.0)
Q3 (2.27-2.64)	146	1.5 (1.3, 1.8)	1.8 (1.6, 2.0)	41.9 (39.0, 45.0)
Q4 (2.65-3.13)	146	1.7 (1.4, 2.0)	1.9 (1.7, 2.2)	45.1 (42.0, 48.4)
Q5 (3.14-7.58)	146	1.9 (1.6, 2.3)	2.1 (1.8, 2.3)	50.3 (46.8, 54.0)
P for trend*		<0.001	0.02	<0.001

\* P for trend of medians in each quintiles

# Saturated fats and CVD: a meta-analysis

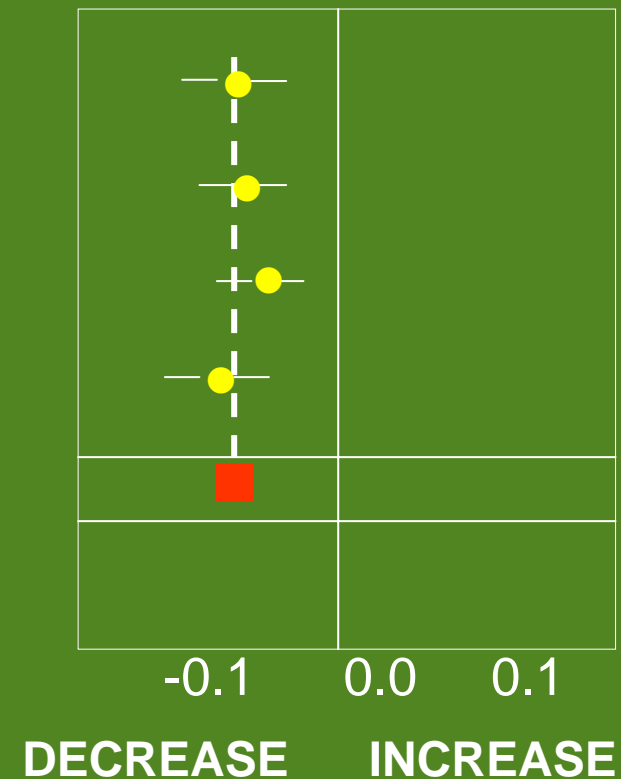


# Saturated fats and CVD: a metanalysis

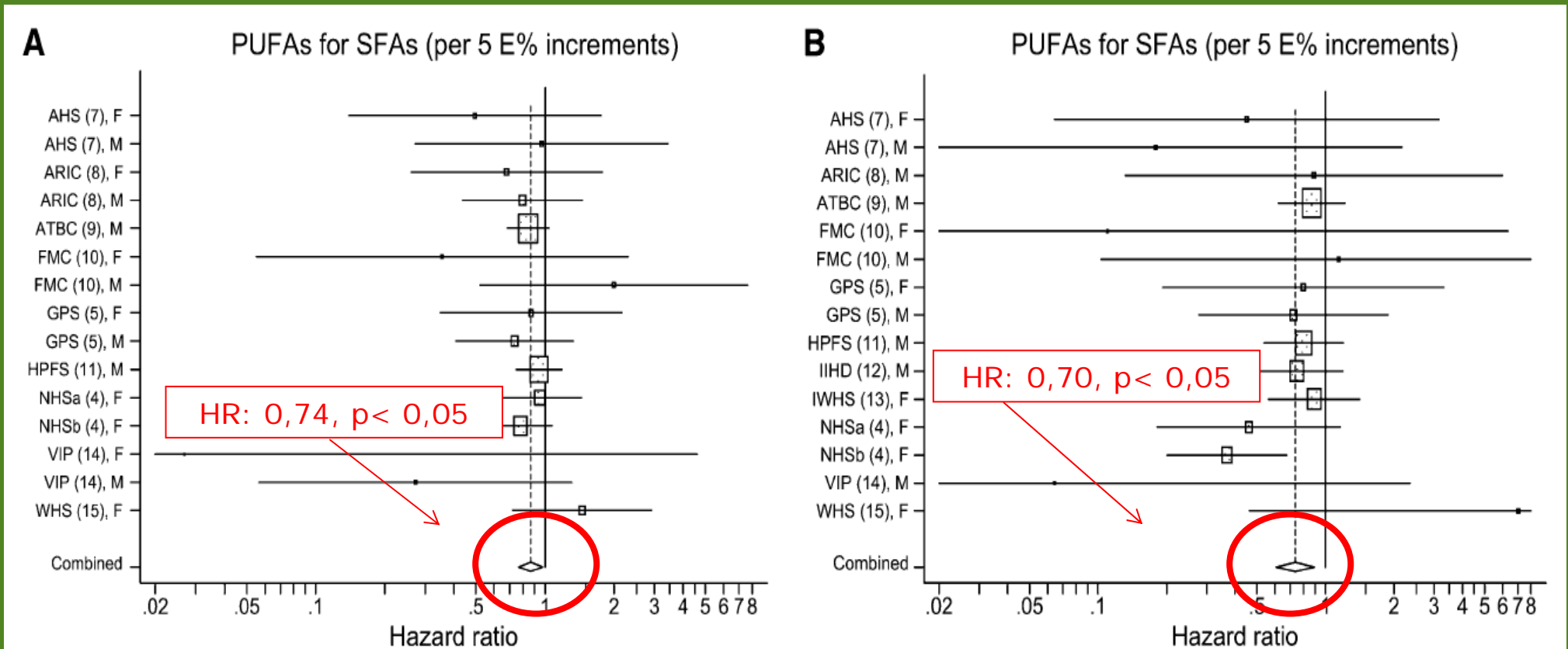


# Effect of different fatty acids on plasma cholesterol in human studies: a meta-analysis

	Multivariate regression coefficient (SE)	Multivariate regression coefficient (99% confidence interv.)
<b>Polyunsaturated fat:</b>		
Crossover design	-0.022 (0.009)	
Parallel square	-0.025 (0.011)	
Latin design	-0.010 (0.008)	
Design	-0.033 (0.005)	
<b>All solid food</b>	<b>-0.026 (0.004)</b>	



# Effetto della sostituzione di SFA con PUFA (5% delle calorie) sulle morti coronariche (A) e sugli eventi CV totali (B)



Jakobsen MU, Am J Clin Nutr 2009

# Major types of dietary fat and risk of coronary heart disease: a pooled analysis of 11 cohort studies

Combined hazard ratios (HRs) for coronary events and coronary deaths per 5% increments in energy intake from polyunsaturated fatty acids (PUFAs) or carbohydrates (CHs) in the Pooling Project of Cohort Studies on Diet and Coronary Disease<sup>1</sup>

	All		Women		Men		<i>P</i> value, test for effect modification by sex
	HR (95% CI)	<i>P</i> value, test for between-studies heterogeneity	HR (95% CI)	<i>P</i> value, test for between-studies heterogeneity	HR (95% CI)	<i>P</i> value, test for between-studies heterogeneity	
Coronary events <sup>2</sup>							
MUFAs for SFAs							
Model 1 <sup>3</sup>	1.39 (1.20, 1.61)		1.33 (1.01, 1.74)		1.47 (1.25, 1.73)		
Model 2 <sup>4</sup>	1.19 (1.00, 1.42)	0.32	1.15 (0.84, 1.58)	0.30	1.23 (0.98, 1.55)	0.32	0.49
PUFAs for SFAs							
Model 1 <sup>3</sup>	0.69 (0.59, 0.81)		0.66 (0.54, 0.81)		0.68 (0.52, 0.87)		
Model 2 <sup>4</sup>	0.87 (0.77, 0.97)	0.70	0.85 (0.68, 1.06)	0.51	0.87 (0.76, 1.01)	0.61	0.84
CHs for SFAs							
Model 1 <sup>3</sup>	1.06 (1.01, 1.12)		0.98 (0.90, 1.06)		1.10 (1.05, 1.16)		
Model 2 <sup>4</sup>	1.07 (1.01, 1.14)	0.51	1.00 (0.89, 1.12)	0.72	1.11 (1.02, 1.20)	0.37	0.13

Jakobsen MU, Am J Clin Nutr 2009

# Major types of dietary fat and risk of coronary heart disease: a pooled analysis of 11 cohort studies

## **Conclusion:**

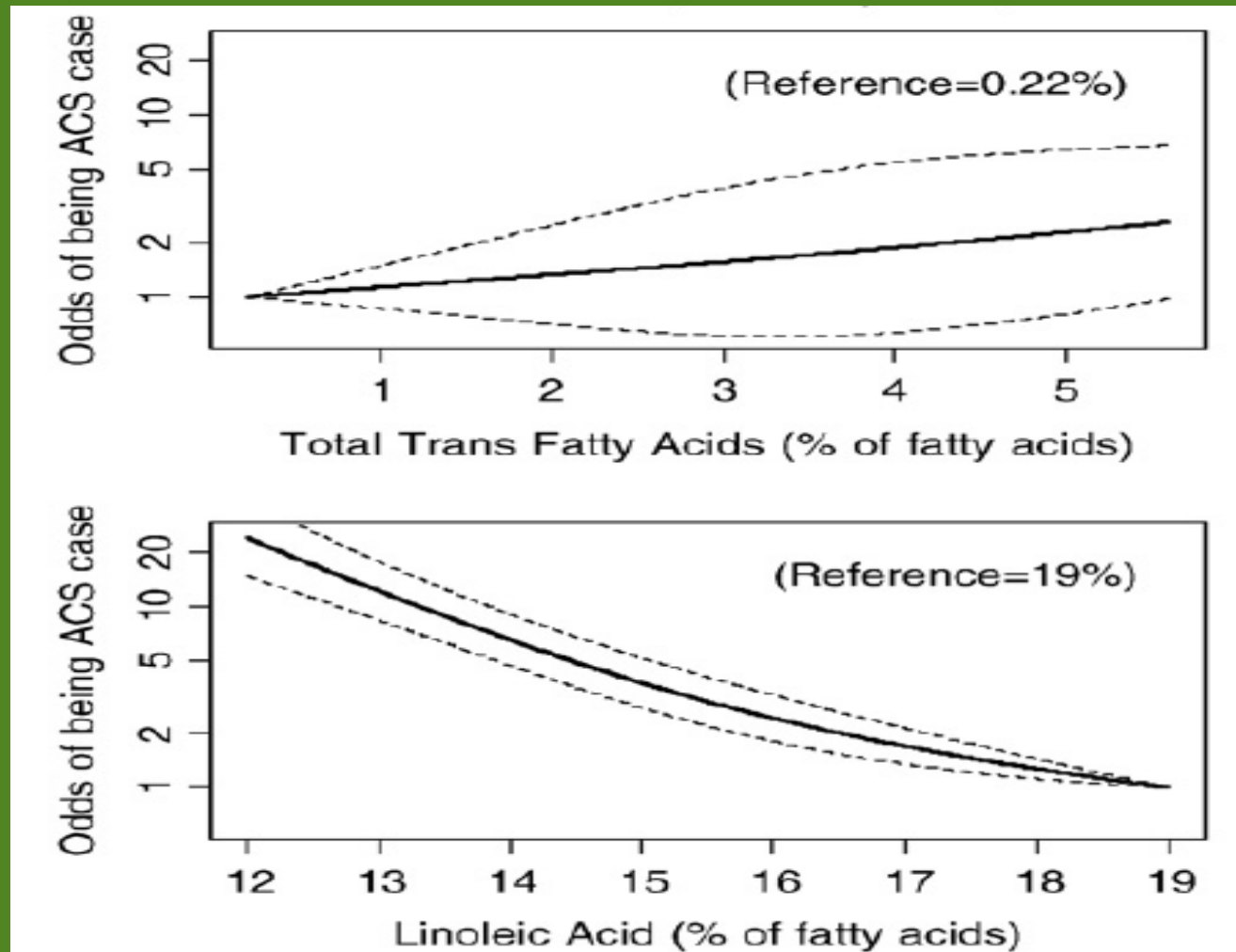
*The associations observed suggest that replacing SFAs with PUFAs rather than MUFAs or carbohydrates prevents CHD over a wide range of intakes.*

## Omega-6 and trans fatty acids in blood cell membranes: A risk factor for acute coronary syndromes?

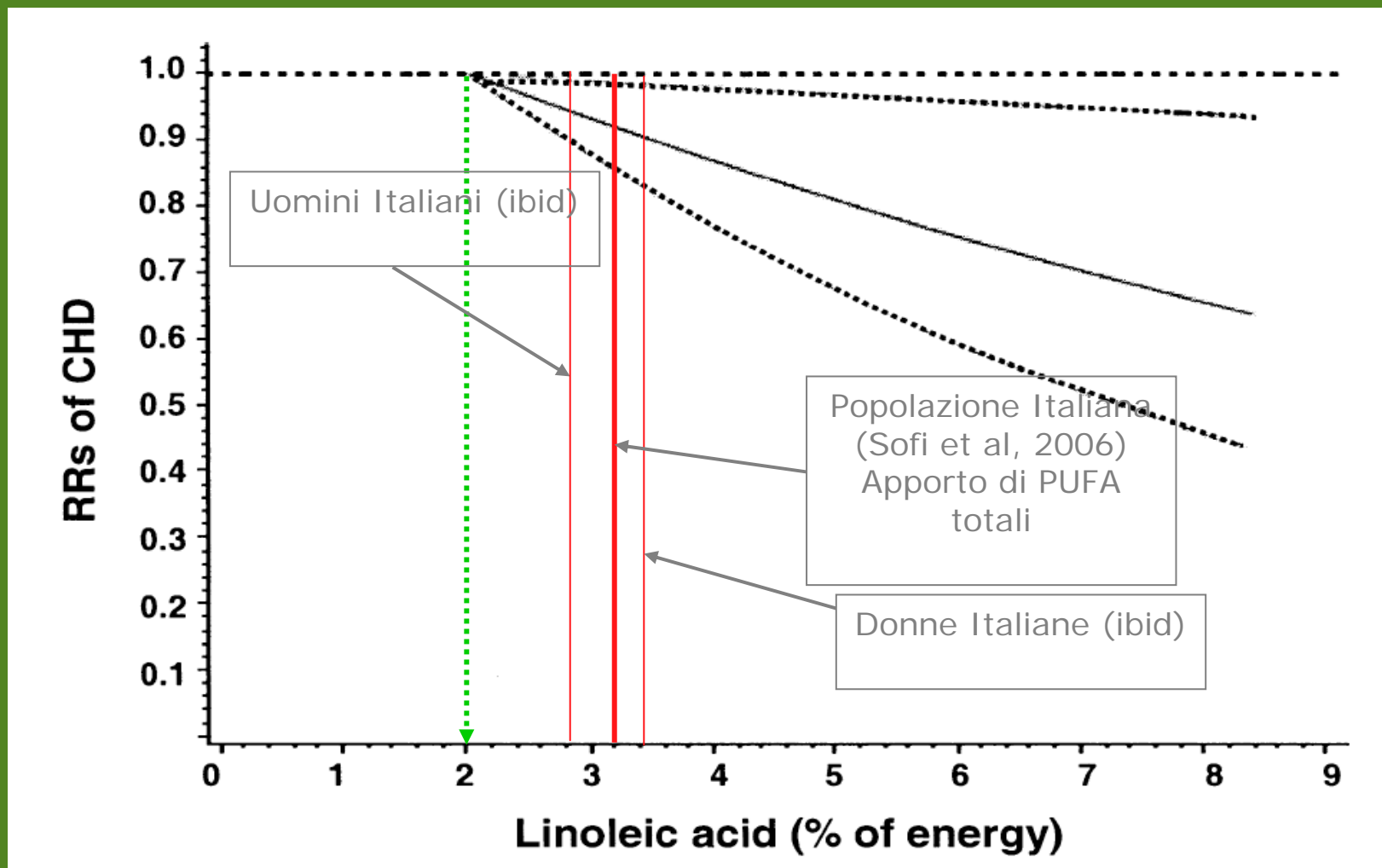
**Table I.** Description of cases and controls

<b>Variable</b>	<b>Cases (n = 768)</b>	<b>Controls (n = 768)</b>	<b>P</b>
Linoleic acid (% total FA)	14.26 ± 2.51	16.45 ± 2.57	<.0001
Arachidonic acid (% total FA)	14.21 ± 3.70	13.72 ± 2.39	.0007
Total <i>trans</i> -fats (% total FA) *	3.78 ± 1.40	3.34 ± 1.29	<.0001
Oleic acid (% total FA)	18.00 ± 3.67	17.29 ± 2.63	<.0001
α-Linoleic acid (% total FA)	0.31 ± 0.17	0.48 ± 0.25	<.0001
Eicosapentaenoic acid (% total FA)	0.46 ± 0.29	0.72 ± 0.53	<.0001
Docosahexaenoic acid (% total FA)	2.93 ± 1.40	3.53 ± 1.57	<.0001

# Omega-6 and trans fatty acids in blood cell membranes: A risk factor for acute coronary syndromes?

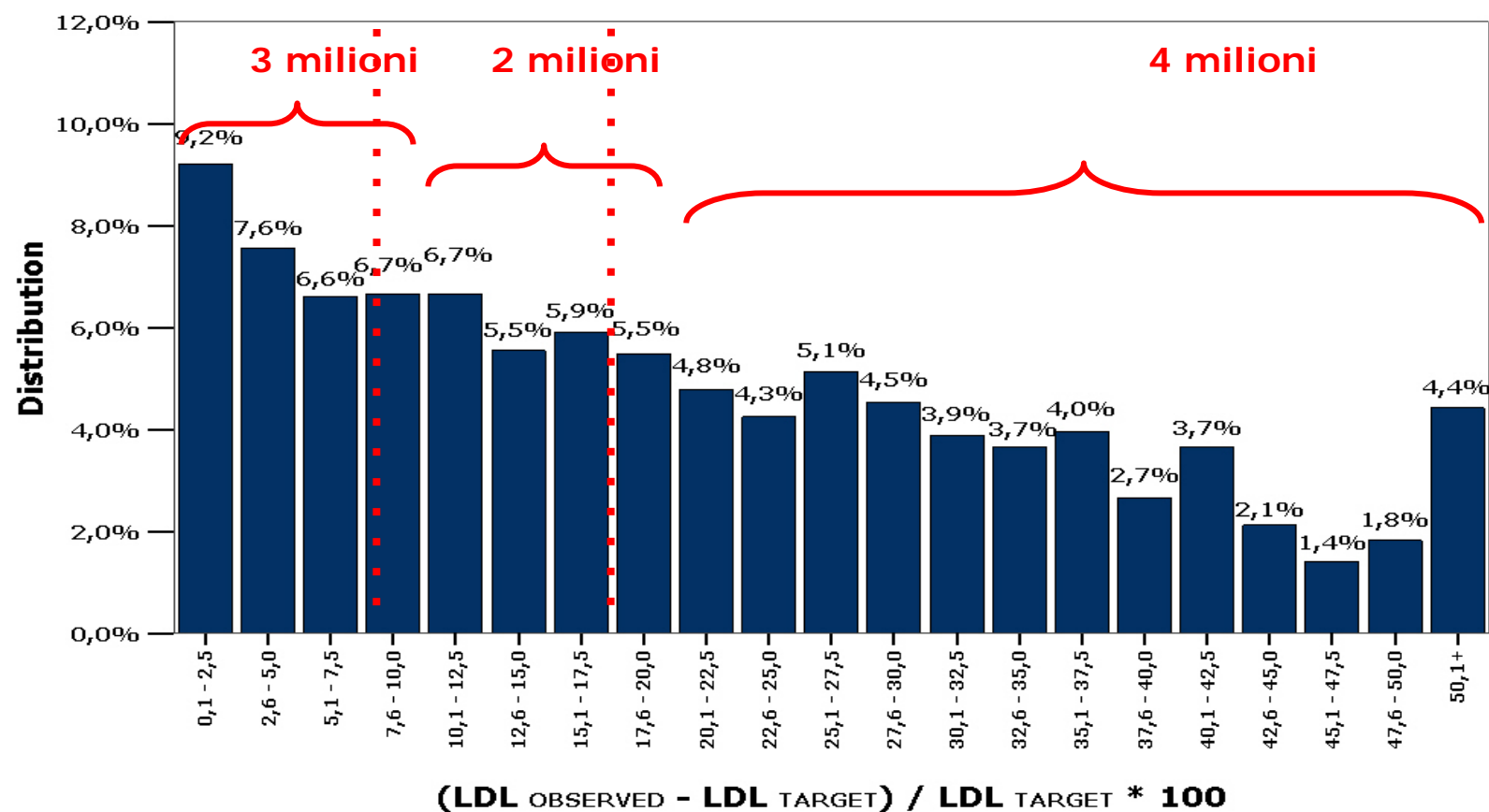


# Rischio di eventi coronarici ed apporto alimentare di acido linoleico nel Nurses' Health Study



# Distanza dal proprio target (%) dei soggetti "non a target" del campione CHECK, e classificazione in gruppi di possibile intervento

**Totale soggetti "non a target" (40-79 anni): 9 milioni**



# Steroli plasmatici utilizzati come indicatori di assorbimento o sintesi del colesterolo

## Markers di sintesi

- Latosterolo
- Colesstenolo
- Desmosterolo
- Squalene

## Markers di assorbimento

- Colestanolo
- Campesterolo
- Sitosterolo
- Avenasterolo

# Markers di assorbimento e sintesi del colesterolo in diverse diete sperimentali

Variables	Diets								<i>P</i>
	HD ( <i>n</i> 29)		LCLF ( <i>n</i> 29)		HCLF ( <i>n</i> 29)		LCHF ( <i>n</i> 18)		
	Mean	SEM	Mean	SEM	Mean	SEM	Mean	SEM	
Total cholesterol§ (mmol/l)	6.01	0.21	5.07*	0.18	5.61*†	0.20	5.55*†	0.27	< 0.001
HDL-cholesterol (mmol/l)	1.41	0.06	1.30	0.05	1.44	0.06	1.41	0.08	0.090
LDL-cholesterol (mmol/l)	3.87	0.22	3.04*	0.17	3.46*†	0.20	3.32*†	0.23	< 0.001
TAG (mmol/l)	1.63	0.16	1.62	0.12	1.58	0.13	1.82	0.27	0.460

HCLF, high-cholesterol low-fat diet; HD, baseline home diet; LCHF, low-cholesterol high-fat diet; LCLF, low-cholesterol low-fat diet.

Variables	HD ( <i>n</i> 29)		LCLF ( <i>n</i> 29)		HCLF ( <i>n</i> 29)		LCHF ( <i>n</i> 18)		<i>P</i>
	Mean	SEM	Mean	SEM	Mean	SEM	Mean	SEM	
Lathosterol§	145	10	134	9	128*	10	236*†‡	22	< 0.001
Cholestanol§	85	6	119*	6	117*	5	133*†	7	< 0.001
Campesterol§	170	14	183	15	169†	15	247*†‡	20	< 0.001
Sitosterol§	113	8	131*	7	119†	8	137*	8	0.041

HCLF, high-cholesterol low-fat diet; HD, baseline home diet; LCHF, low-cholesterol high-fat diet; LCLF, low-cholesterol low-fat diet.

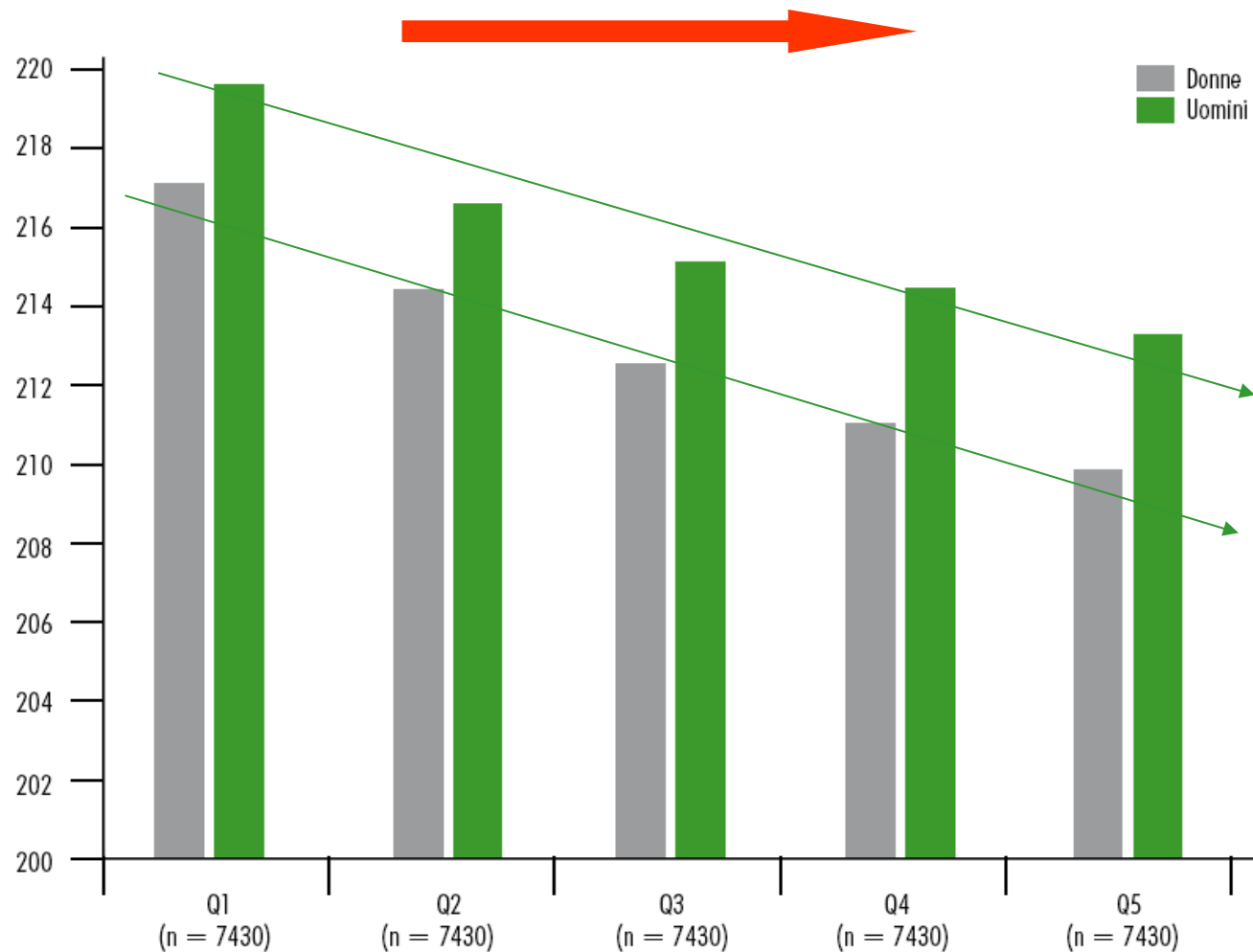
a: ratio su Chol

Nissinen MJ et al. Brit J Nutr 2008

# Effects of a phytosterol-enriched dairy product on lipids, sterols and 8-isoprostane in hypercholesterolemic patients: A multicenter Italian study

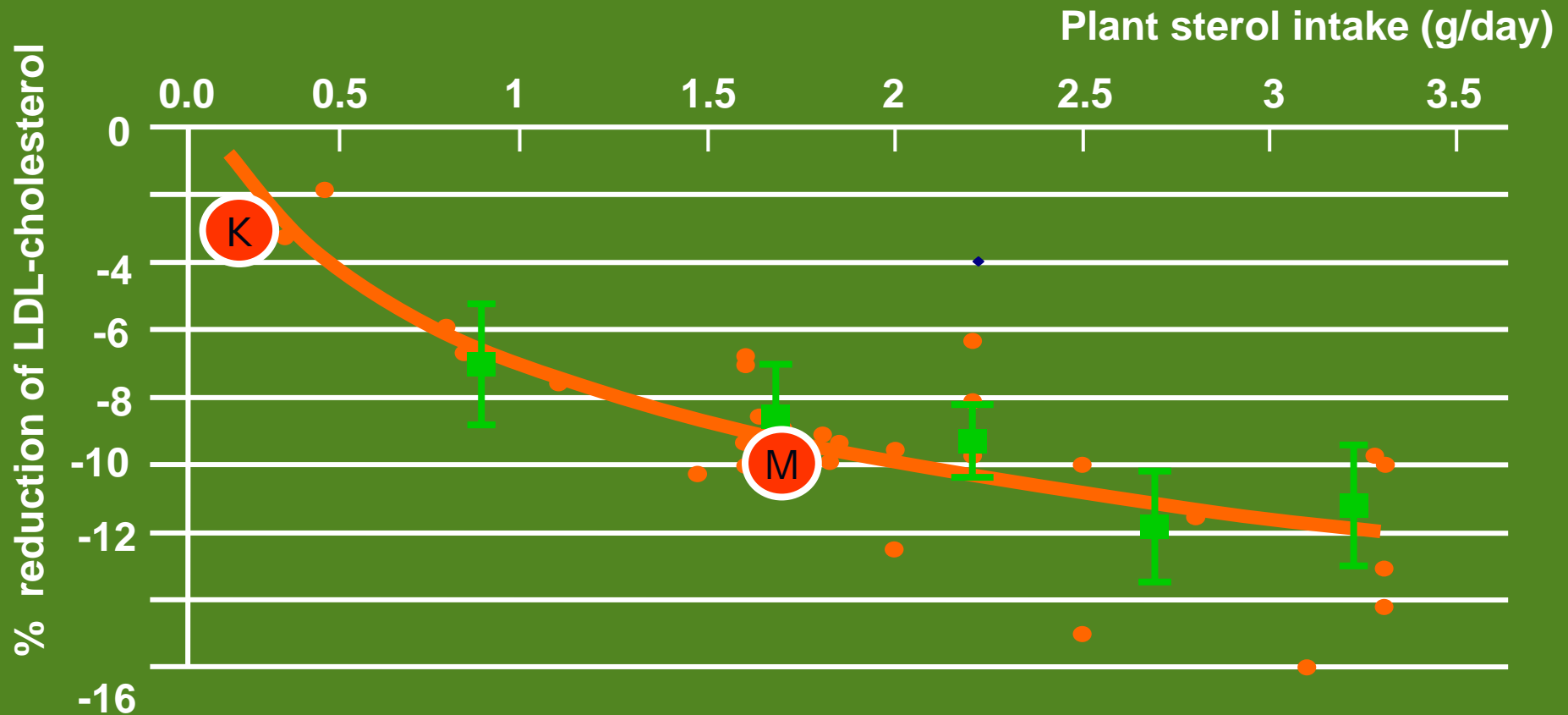
	PS-enriched fermented milk <i>N</i> = 60	Control fermented milk <i>N</i> = 56
<i>Total cholesterol mg/dl</i>		
Baseline	263.5 ± 2.6	260.0 ± 3.2
3 weeks	226.9 ± 3.3*	242.5 ± 3.5
6 weeks	231.0 ± 3.2*	243.1 ± 4.2
<i>Triglycerides mg/dl</i>		
Baseline	126.8 ± 6.8	125.4 ± 7.1
3 weeks	117.0 ± 5.5	125.6 ± 7.0
6 weeks	131.6 ± 9.1	128.5 ± 7.6
<i>LDL cholesterol mg/dl</i>		
Baseline	166.2 ± 2.0	163.7 ± 2.1
3 weeks	148.7 ± 3.1*	160.1 ± 2.8
6 weeks	147.4 ± 2.8*	160.5 ± 3.1
<i>HDL cholesterol, mg/dl</i>		
Baseline	51.6 ± 1.9	50.7 ± 1.9
3 weeks	51.9 ± 1.9	52.7 ± 2.1
6 weeks	53.4 ± 2.2	52.7 ± 2.1

# Fitosteroli naturalmente contenuti negli alimenti e colesterolemia: uno studio epidemiologico scandinavo



Colesterolemia totale nei differenti quintili di apporto alimentare di fitosteroli.

# Cholesterol lowering with plant sterols in fat-based foods: dose-response relationship



- data of ~ 30 placebo-controlled initiated studies with phytosterol-enriched spreads
- data (mean plus 95% confidence interval) from meta-analysis of 41 studies with phytosterols or stanols (Katan et al, Mayo Clin Proc. 2003)

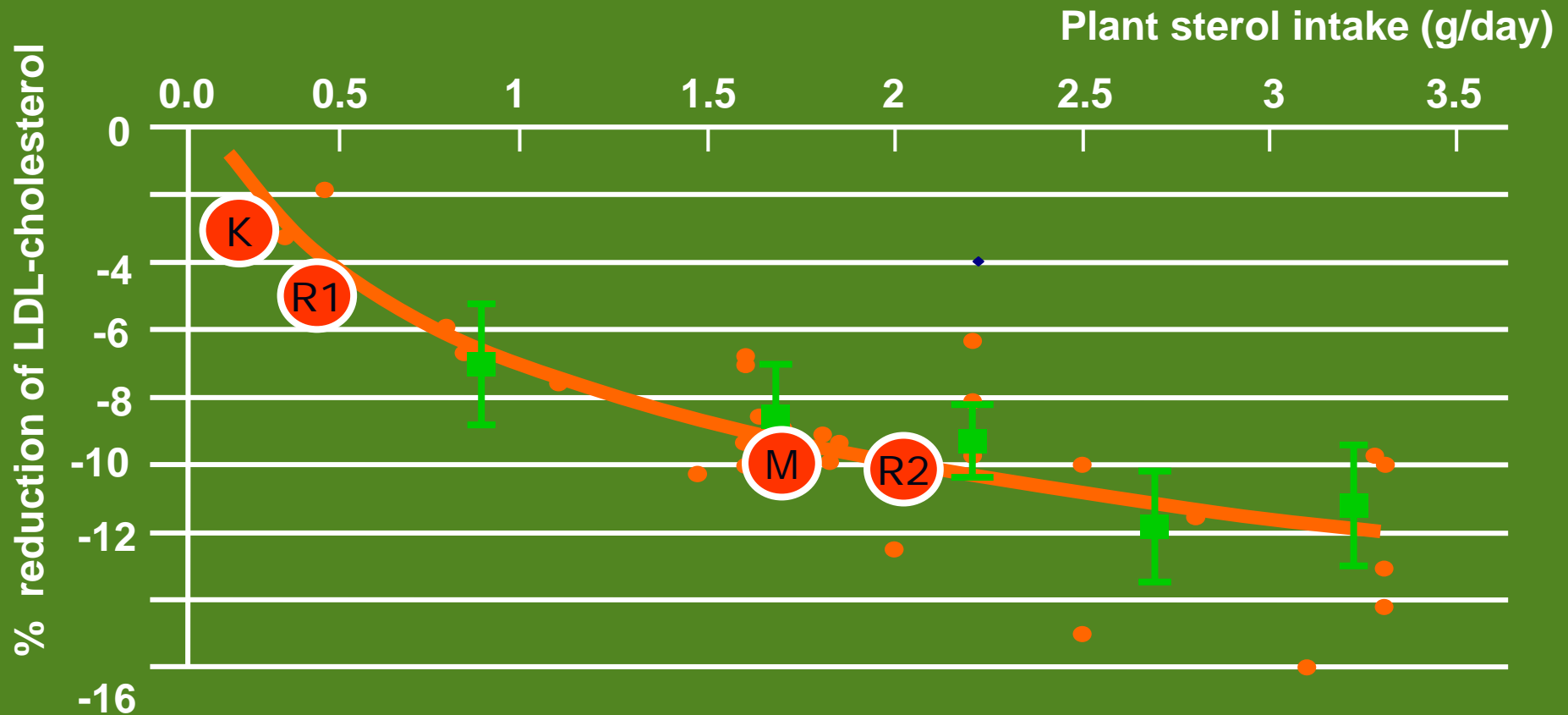
# Dose effects of dietary phytosterols on cholesterol metabolism: a controlled study

	Phytosterol intake		
	59 mg/d (control)	459 mg/d (moderate)	2059 mg/d (high)
<b>Cholesterol absorption</b>			
Percentage cholesterol absorbed (%)	69.9 ± 2.1	62.8 ± 2.1 <sup>2</sup>	52.7 ± 2.1 <sup>2,3</sup>
Dietary cholesterol absorbed (mg/d)	139 ± 8	121 ± 8 <sup>4</sup>	105 ± 8 <sup>2,5</sup>
Biliary cholesterol absorbed (mg/d)	1418 ± 167	1390 ± 167	1244 ± 167
<b>Plasma sterol ratios</b>			
Phytosterols/total cholesterol (µg/mg)	7.4 ± 1.3	14.1 ± 1.3 <sup>2</sup>	25.8 ± 1.3 <sup>2,3</sup>
Cholestanol/total cholesterol (µg/mg) <sup>6</sup>	1.09 ± 0.08	0.95 ± 0.08 <sup>2</sup>	0.85 ± 0.08 <sup>2,5</sup>
Lathosterol/total cholesterol (µg/mg) <sup>7</sup>	1.22 ± 0.13	1.51 ± 0.14 <sup>2</sup>	1.71 ± 0.14 <sup>2,3</sup>
<b>Serum lipid concentrations</b>			
Total cholesterol (mg/dL)	211 ± 6	204 ± 6	198 ± 6 <sup>2</sup>
LDL cholesterol (mg/dL)	139 ± 4	132 ± 4	126 ± 4 <sup>2</sup>
HDL cholesterol (mg/dL)	50 ± 3	50 ± 3	51 ± 3
Triglycerides (mg/dL)	112 ± 9	112 ± 9	104 ± 9
Non-HDL cholesterol (mg/dL)	161 ± 5	154 ± 5	146 ± 5 <sup>2</sup>
LDL cholesterol/HDL cholesterol	2.92 ± 0.16	2.75 ± 0.16 <sup>4</sup>	2.58 ± 0.16 <sup>2,5</sup>

# Dose effects of dietary phytosterols on cholesterol metabolism: a controlled study

	Phytosterol intake		
	59 mg/d (control)	459 mg/d (moderate)	2059 mg/d (high)
<b>Cholesterol absorption</b>			
Percentage cholesterol absorbed (%)	69.9 ± 2.1	62.8 ± 2.1 <sup>2</sup>	52.7 ± 2.1 <sup>2,3</sup>
Dietary cholesterol absorbed (mg/d)	139 ± 8	121 ± 8 <sup>4</sup>	105 ± 8 <sup>2,5</sup>
Biliary cholesterol absorbed (mg/d)	1418 ± 167	1390 ± 167	1244 ± 167
<b>Plasma sterol ratios</b>			
Phytosterols/total cholesterol (μg/mg)	7.4 ± 1.3	14.1 ± 1.3 <sup>2</sup>	25.8 ± 1.3 <sup>2,3</sup>
Cholestanol/total cholesterol (μg/mg) <sup>6</sup>	1.09 ± 0.08	0.95 ± 0.08 <sup>2</sup>	0.85 ± 0.08 <sup>2,5</sup>
Lathosterol/total cholesterol (μg/mg) <sup>7</sup>	1.22 ± 0.13	1.51 ± 0.14 <sup>2</sup>	1.71 ± 0.14 <sup>2,3</sup>
<b>Serum lipid concentrations</b>			
Total cholesterol (mg/dL)	211 ± 6	204 ± 6	198 ± 6 <sup>2</sup>
LDL cholesterol (mg/dL)	139 ± 4	132 ± 4	126 ± 4 <sup>2</sup>
HDL cholesterol (mg/dL)	50 ± 3	50 ± 3	51 ± 3
Triglycerides (mg/dL)	112 ± 9	112 ± 9	104 ± 9
Non-HDL cholesterol (mg/dL)	161 ± 5	154 ± 5	146 ± 5 <sup>2</sup>
LDL cholesterol/HDL cholesterol	2.92 ± 0.16	2.75 ± 0.16 <sup>4</sup>	2.58 ± 0.16 <sup>2,5</sup>

# Cholesterol lowering with plant sterols in fat-based foods: dose-response relationship



- data of ~ 30 placebo-controlled initiated studies with phytosterol-enriched spreads
- data (mean plus 95% confidence interval) from meta-analysis of 41 studies with phytosterols or stanols (Katan et al, Mayo Clin Proc. 2003)

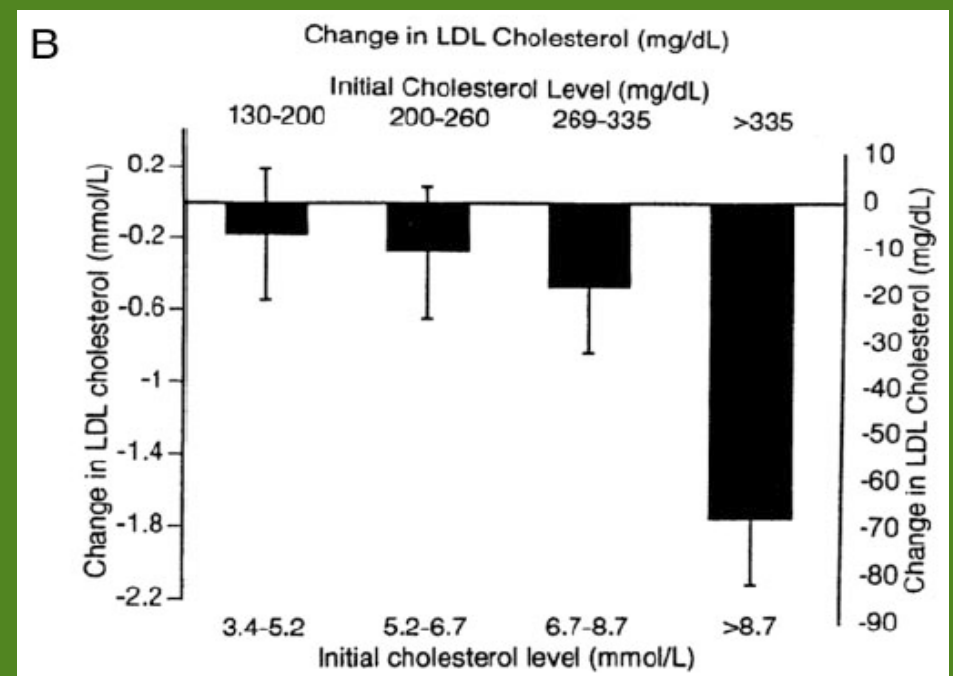
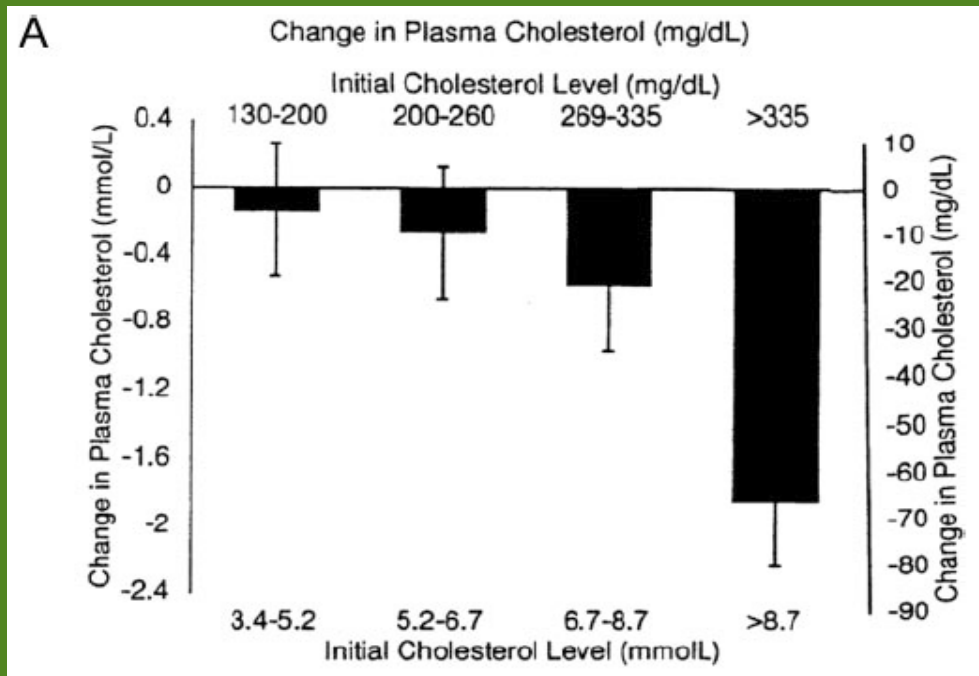
# Mean serum concentrations of total and LDL cholesterol following a 3 week treatment with 5 g/day of oat beta-glucan

<b>Total cholesterol</b>	<b>Control group</b>	<b>Beta-glucan group</b>	<b>p</b>
Run-in period (mmol/L)	5.57 ± 0.66	5.94 ± 0.76	
Test period (mmol/L)	5.67 ± 0.66	5.75 ± 0.58	
Change (mmol/L)	0.11 ± 0.37	-0.19 ± 0.39	0.012
Change (%)	2.14 ± 6.81	-2.70 ± 6.06	
<b>LDL cholesterol</b>			
Run-in period (mmol/L)	3.45 ± 0.68	3.87 ± 0.80	
Test period (mmol/L)	3.54 ± 0.71	3.67 ± 0.61	
Change (mmol/L)	0.10 ± 0.34	-0.21 ± 0.37	0.005
Change (%)	3.23 ± 9.99	-4.51 ± 8.50	

# La nuova normativa EFSA sul beta-glucano

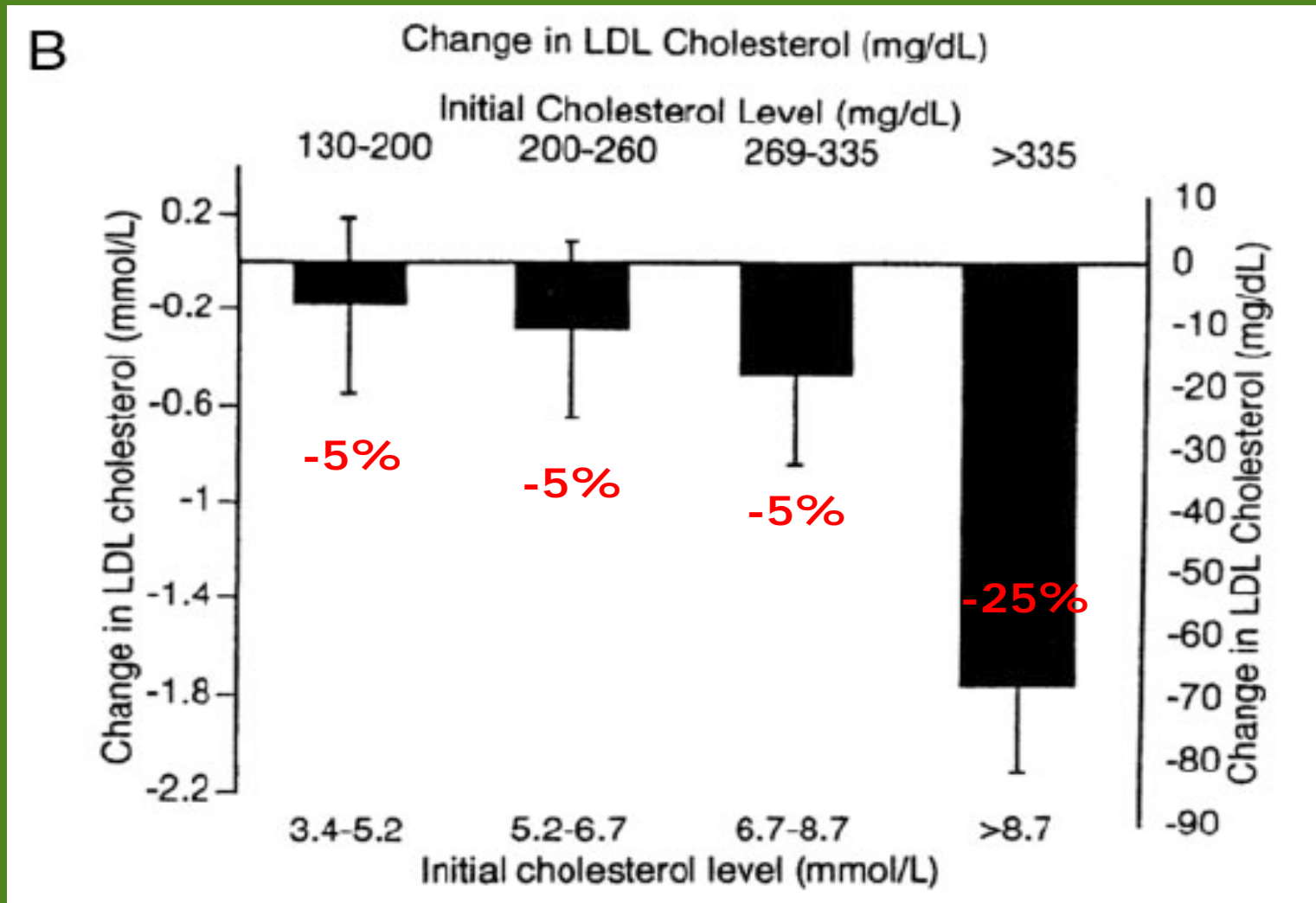
- La dose giornaliera efficace è fissata in almeno 3 grammi
- Il prodotto deve essere in grado di fornire l'intera dose giornaliera
- Il wording suggerito è "aiuta a mantenere normali valori della colesterolemia"
- La normativa non è ancora entrata in vigore

# Changes in Tot Chol (A) and LDL-c (B) and initial cholesterol levels in a meta-analysis of clinical trials with soy protein.



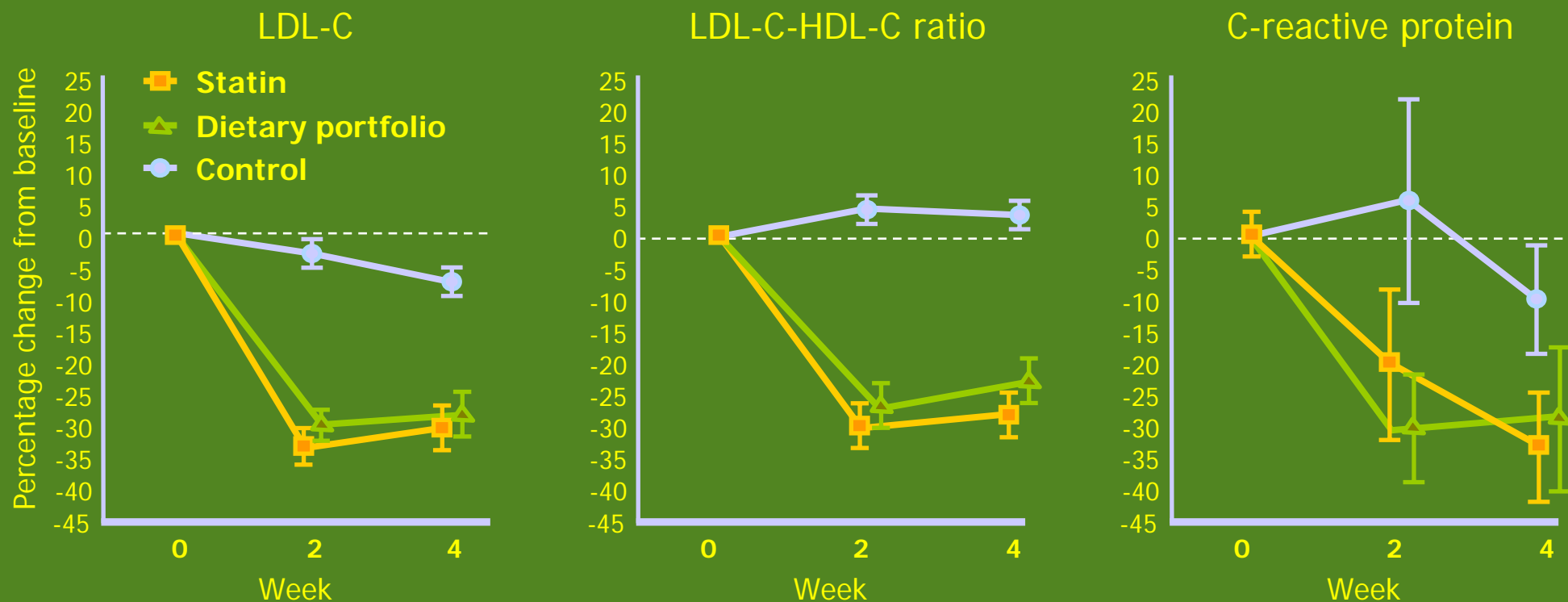
Anderson JW et al, N Engl J Med 1995

# Hypocholesterolaemic effects of soy proteins: effect of basal cholesterol levels



Anderson JW et al, N Engl J Med 1995

# Effects of a "Dietary Portfolio" vs Lovastatin on Serum Lipid and CRP



.density lipoprotein cholesterol-high=C-HDL ;density lipoprotein Cholesterol-low=C-LDL  
 approximately twice the SE represents ,with the number of participants involved ,because )SE(Values are expressed as mean  
 a significant difference

# Assessment of the long term effects of a dietary portfolio of cholesterol-lowering foods in patients with hypercholesterolemia

Blood lipid measurements at baseline and after 1 y of the ad libitum dietary portfolio study in 66 participants<sup>1</sup>

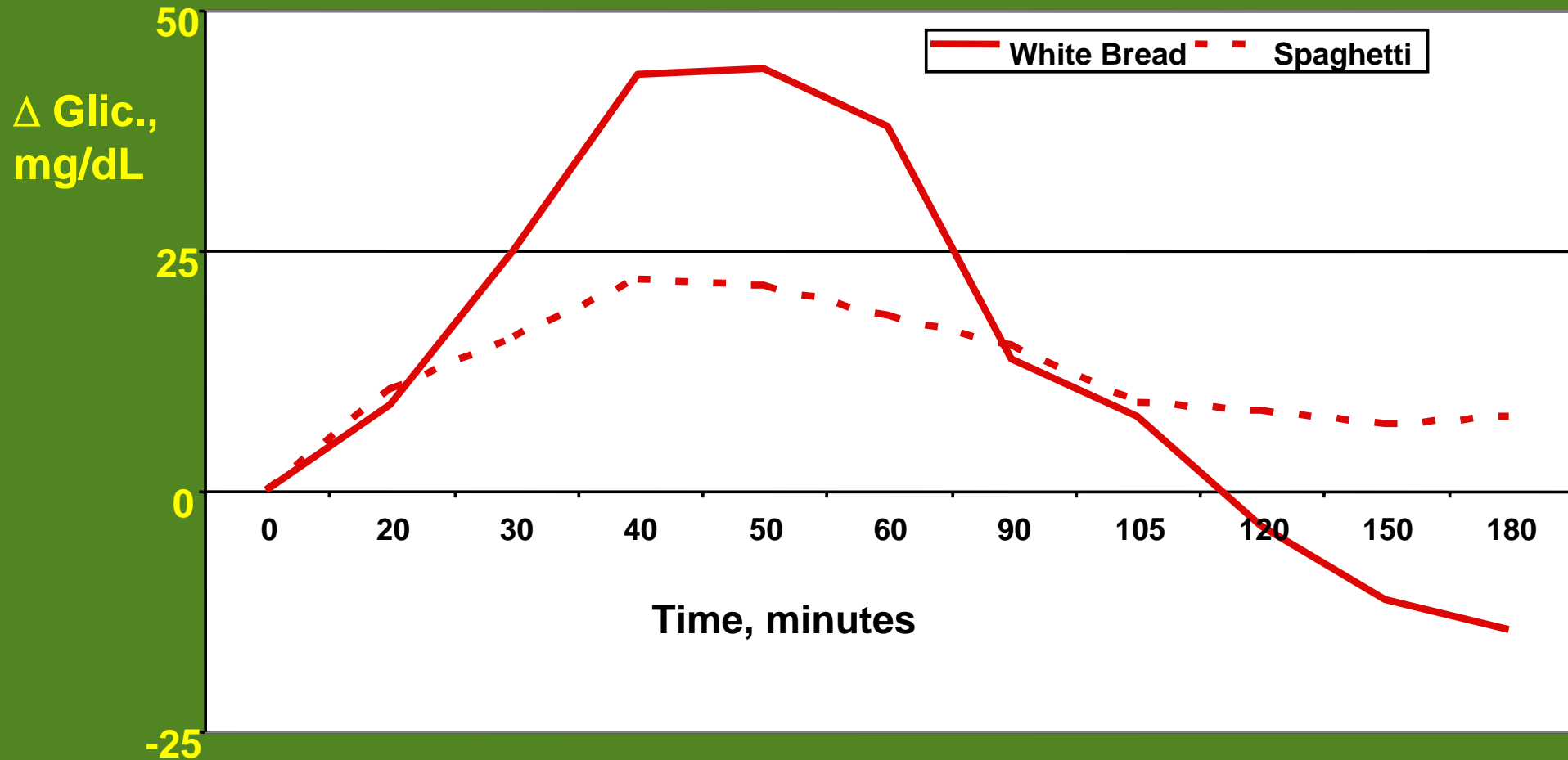
	Total C	LDL-C	HDL-C	Triacylglycerol	Total:HDL-C	LDL-C:HDL-C
Prestudy	6.75 ± 0.10	4.48 ± 0.09	1.24 ± 0.04	2.29 ± 0.14	5.76 ± 0.16	3.82 ± 0.12
1 y	6.05 ± 0.11	3.87 ± 0.10	1.28 ± 0.05	1.98 ± 0.11	5.03 ± 0.16	3.23 ± 0.12
Change	-0.70 ± 0.09	-0.61 ± 0.09	0.04 ± 0.02	-0.31 ± 0.11	-0.73 ± 0.10	-0.59 ± 0.08
<i>p</i> <sup>2</sup>	< 0.0001	< 0.0001	0.0256	0.0051	< 0.0001	< 0.0001

<sup>1</sup> All values are  $\bar{x} \pm SE$ . Total C, total cholesterol; LDL-C, LDL cholesterol; HDL-C, HDL cholesterol. A last-observation-carried-forward analysis was performed for 11 participants for weeks 0 (*n* = 1), 2 (*n* = 1), 12 (*n* = 3), 24 (*n* = 2), 32 (*n* = 3), and 42 (*n* = 1).

<sup>2</sup> Significance of change was determined with Student's paired *t* test (2-tailed, PROC MEANS) (28).

**-15%**

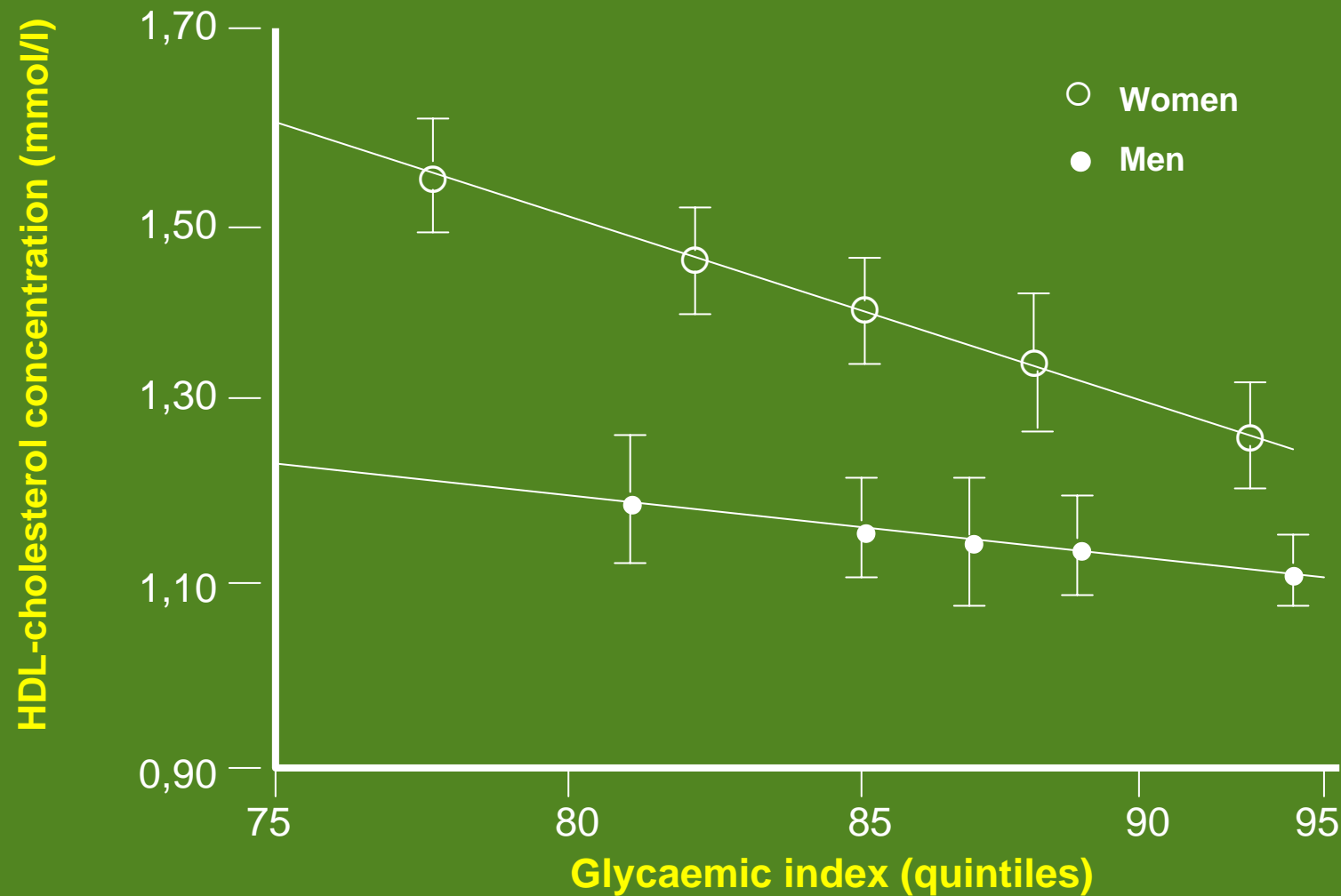
# GLYCEMIC RESPONSE AFTER A WHITE BREAD OR A SPAGHETTI MEAL



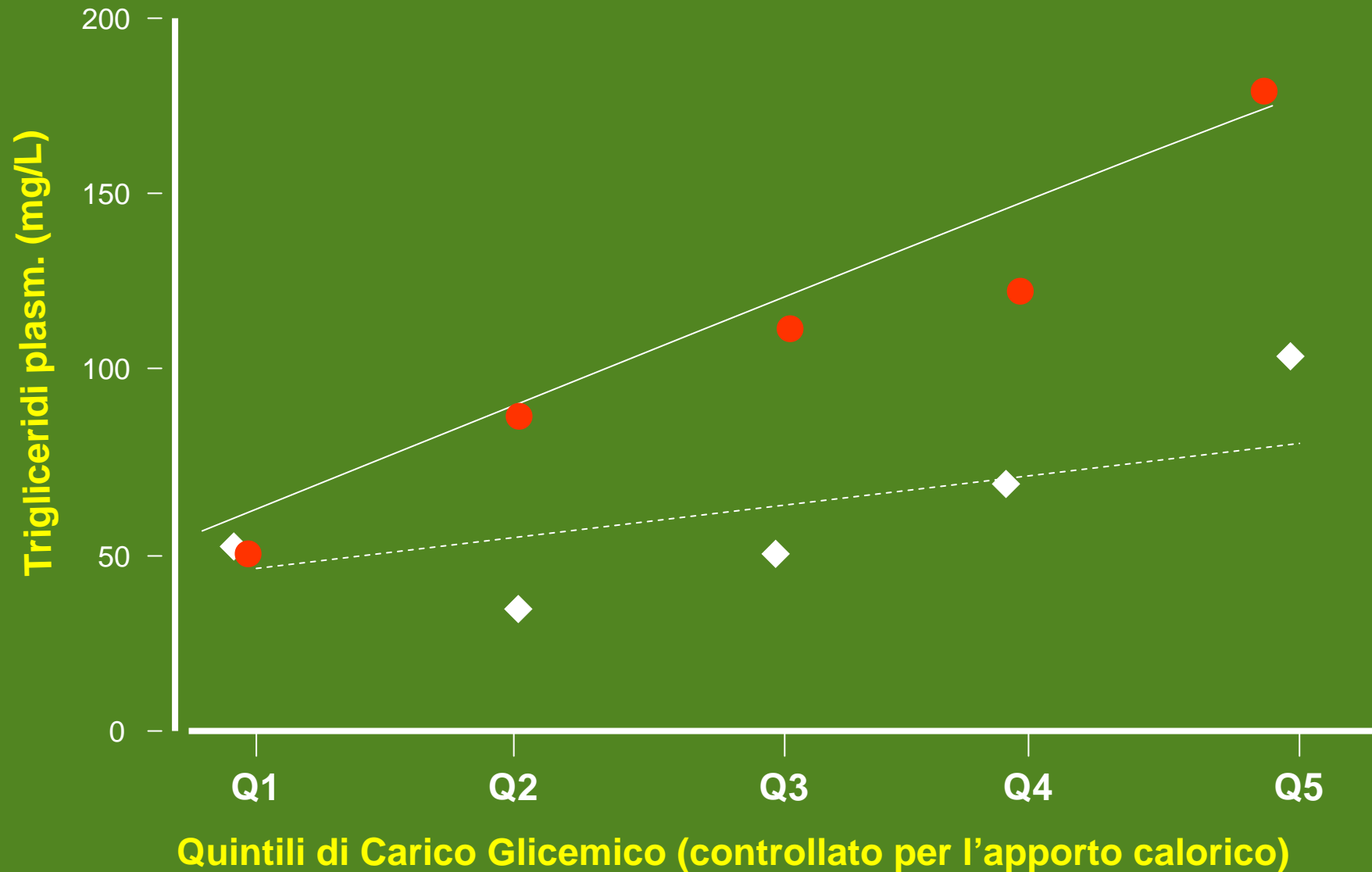
# Indice Glicemico (IG), relativo al Pane Bianco, di alcuni alimenti

Cibo	Indice glicemico (%)
Pane bianco	100
Pomodori	13
Ciliegie	32
Fagioli	40-60
Mele	52
Pasta (spaghetti)	52
Pasta (maccheroni)	68
Pizza	86
Saccarosio	92
Polenta	106
Cornflakes	100-120
Miele	120
Patate bollite	120
Glucosio	138

# Relation Between HDL-Cholesterol Concentration and Glycaemic Index of the Diet in Men and Women



## Carico Glicemico medio della dieta e trigliceridemia in donne con (●) e senza (◆) sovrappeso (BMI >25)



# Effetto dell'attività fisica sul profilo dei lipidi plasmatici: lo studio "Heritage"

- Soggetti studiati: 200 uomini, di età <65 anni, sani e sedentari
- Training: 60 sessioni di training aerobico in 21 settimane (da 1 a 4 sessioni per settimana)
- Effetto del training sul profilo lipidico:

	$\Delta$ HDL2c	$\Delta$ TG	$\Delta$ TC/HDLc
Normolipemici	+ 3 %	+ 2 %	- 2 %
TG alti	+ 9 %	- 13 %	- 4 %
HDL basse, TG alti	+ 18 %	- 17 %	- 9 %

Le variazioni indotte da un training di tipo aerobico sono tanto maggiori quanto meno favorevoli sono le condizioni di partenza del profilo lipidico.