## **Canola Oil and Heart Health**



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#### Richardson Centre for Functional Foods and Nutraceuticals University of Manitoba

#### Diabetes Care and Education Practice Group 18 June 2010 Webinar







# **Canola Oil**



- Canola oil comes from the crushed seeds of canola plants
- Canola is its own plant species, different from rapeseed
- > Developed by traditional cross-breeding in the late 1950s/1960s to remove undesirable components of rapeseed (erucic acid and glucosinolates)
- > Approved for sale in Canada in 1974 and in the U.S. in 1985 (GRAS)
- **FDA authorized qualified health claim for canola oil in October 2006**
- Important economic crop in Canada as world's largest exporter of canola seed, oil and meal
- U.S. #1 importer of canola oil and meal from Canada, also grows its own canola
- Canola oil is world's third leading vegetable oil by volume:
  - #1 oil consumed in Canada, #2 in U.S.



## **Composition of Canola Oil**

High in monounsaturated fat **Canola Oil** High in omega-3 fat High in plant sterols Antioxidant vitamin E Low in saturated fat

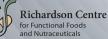
Other health-promoting constituents



## **Comparison of Dietary Fats & Oils**

#### **DIETARY FAT**

Canola oil	7 21	11			61
Safflower oil	8 14	1			77
Flaxseed oil	<mark>9 16</mark>			57	18
Sunflower oil	<b>12 71</b>			1	16
Corn oil	13 57			1	29
Olive oil	15 9	1			75
Soybean oil	15 54			8	23
Peanut oil	19	33	*		48
Cottonseed oil	27	54		*	23 48 19 47 39 28 28 2
Lard	43		9 1		47
Palm oil	51		10 *		39
Butter	68			3 1	28
Coconut oil	91				2 7
	SATURATED FAT	POLYUNSATU	RATED FAT	MONOUNSAT	URATED FAT
		<b>linoleic a</b> (an omega	r <b>cid alpha-linol</b> a-6 fatty acid) (an omega-3	l <b>enic acid</b> oleic acid 3 fatty acid) (an omega-	9 fatty acid)
			*Trace	Fatty acid content normal	ized to 100%



www.canolainfo.org

## **Dietary Fat Intakes & Recommendations**

	CURRENT INTAKES		RECOMMENDED INTAKE		
FAT (% of daily energy)	MEDITERRANEAN <sup>1</sup>	UNITED STATES <sup>2</sup>	ADA/DC (Dietary Guidelines 2005) <sup>3</sup>	AHA NCEP (ATP III) <sup>4</sup>	
Total Fat	33-40%	33%	20-35%	25-35%*	
Saturated	≤8%	11%	< 10%	< 7%	
Monounsaturated	16-29%	12%	< 25%	≤ 20%	
Polyunsaturated	< 7%	< 7%	< 10%	≤ 10%	
linoleic acid		14.7 g/d	5-10% (12-17g/d)	5-10%	
α-linoleinc acid		1.5 g/d	0.6-1.2% (1.1-1.6 g/d)	1.5-3.0 g/d	
n-6:n-3 ratio		9.8:1	4:1		
EPA+DHA		100-130 mg/d	500 mg/d	900 mg/d	

\* The 25–35% fat recommendation allows for increased intake of unsaturated fat in place of carbohydrates in people with the metabolic syndrome or diabetes

Note: n-6:n-3 ratio decreased from 12.4:1 to 10.6:1 from 1985 to 1994, reflected by a ~5.5-fold increase in canola oil use<sup>5</sup>

Richardson Centre <sup>1</sup>Perez-Jimenez et al., *Atherosclerosis* 2002;163:385-98; <sup>2</sup>Means of US male and females (ages 20-59) from the NHANES, 1999for Functional Foods 2002; <sup>3</sup>ADA Reports. *J Am Diet Assoc* 2007;107:1599-1611; <sup>4</sup>JAMA 2001 May 16;285:2486-97; <sup>5</sup>Kris-Etherton et al., *Am J Clin Nutr* 2000;71:179**Current Research** 

Dietary Modeling Shows that the Substitution of Canola Oil for Fats Commonly Used in the United States Would Increase Compliance with Dietary Recommendations for Fatty Acids

GUY H. JOHNSON, PhD; DEBRA R. KEAST, PhD; PENNY M. KRIS-ETHERTON, PhD, RD

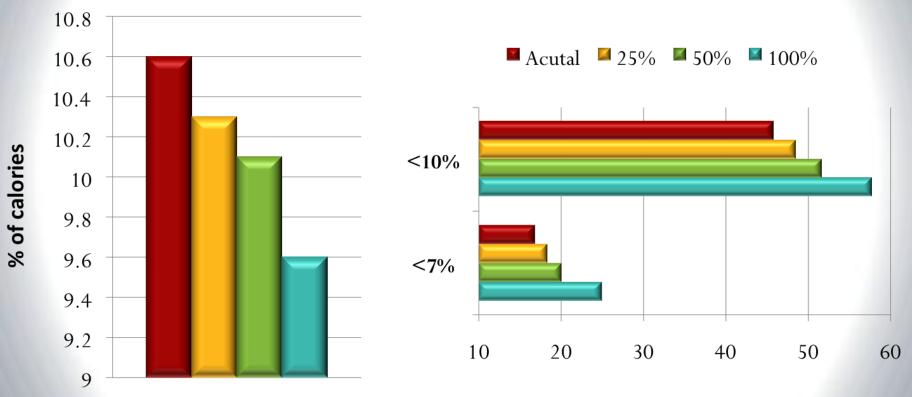
**Objective:** "To examine the effect of substituting canola oil for selected vegetable oils and canola oil-based margarine for other spreads on energy, fatty acid, and cholesterol intakes among US adults".

Design: Adults aged ≥ 20 years (n=8,983) from the 1999-2002
National Health and Nutrition Examination Survey (NHANES)
25%, 50%, 100% substitution of canola oil and canola oil-based margarine for dietary corn, cottonseed, safflower, soybean, and other vegetable oils and spreads.

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Johnson et al., J Am Diet Assoc 2007;107:1726-34

#### Substitution of Canola Oil for Other Oils; Effect on SFA Intakes and Percent of People Meeting Current Dietary Recommendations

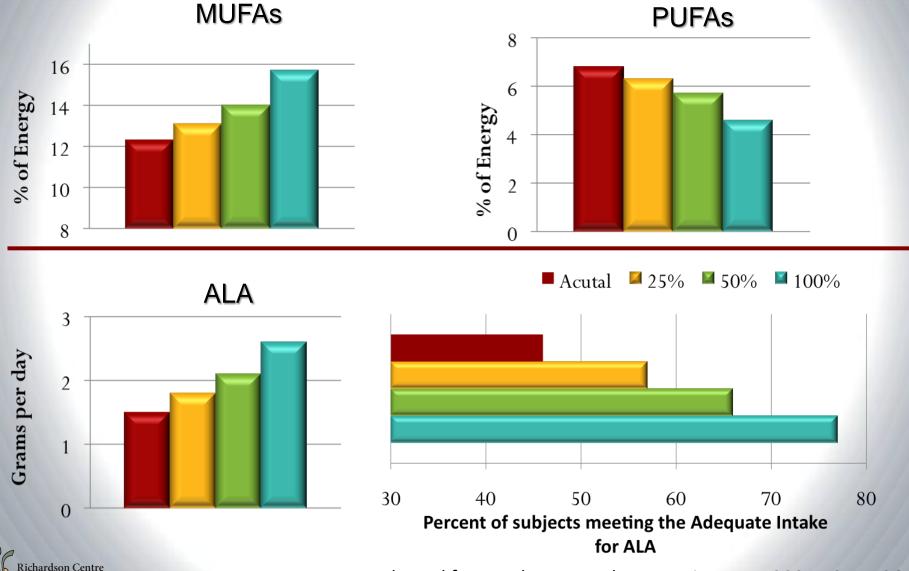


Percent of subjects meeting SFA recommendations



Adapted from: Johnson et al., J Am Diet Assoc 2007;107:1726-34

#### Substitution of Canola Oil for Other Oils; Effect on MUFA, PUFA, and ALA Intakes



for Functional Foods and Nutraceuticals Adapted from: Johnson et al., J Am Diet Assoc 2007;107:1726-34

**Current Research** 

Dietary Modeling Shows that the Substitution of Canola Oil for Fats Commonly Used in the United States Would Increase Compliance with Dietary Recommendations for Fatty Acids

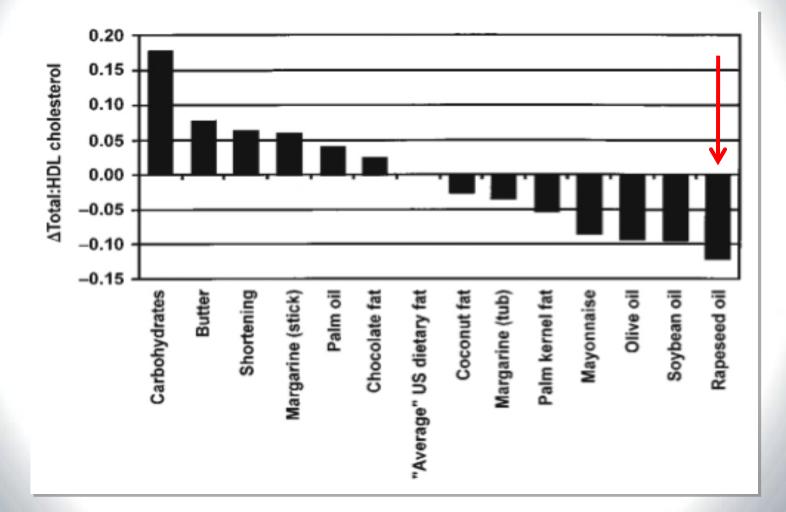
GUY H. JOHNSON, PhD; DEBRA R. KEAST, PhD; PENNY M. KRIS-ETHERTON, PhD, RD

**Conclusion:** "Substitution of canola oil and canola oil-based margarine for most other vegetable oils and spreads increased compliance with dietary recommendations for saturated fatty acid, monounsaturated fatty acid, and  $\alpha$ -linolenic acid, but not for linoleic acid, among US adults".



Johnson et al., J Am Diet Assoc 2007;107:1726-34

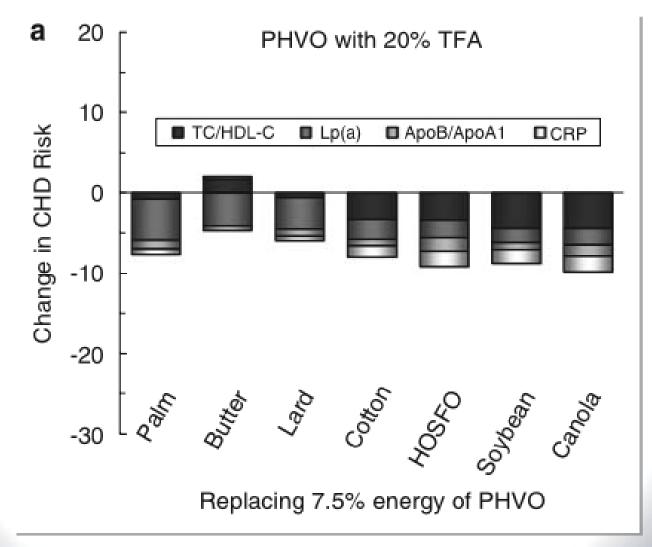
#### **Isoenergetic Replacement of 10% Energy from Mixed Fat in the Average American Diet with a Specific Fat or CHO**





Mensink et al., Am J Clin Nutr 2003;77:1146-55

#### Isoenergetic Replacement of 7.5% Energy from Partially Hydrogenated Vegetable Oil (20% TFAs) with Alternative Fats or Oils





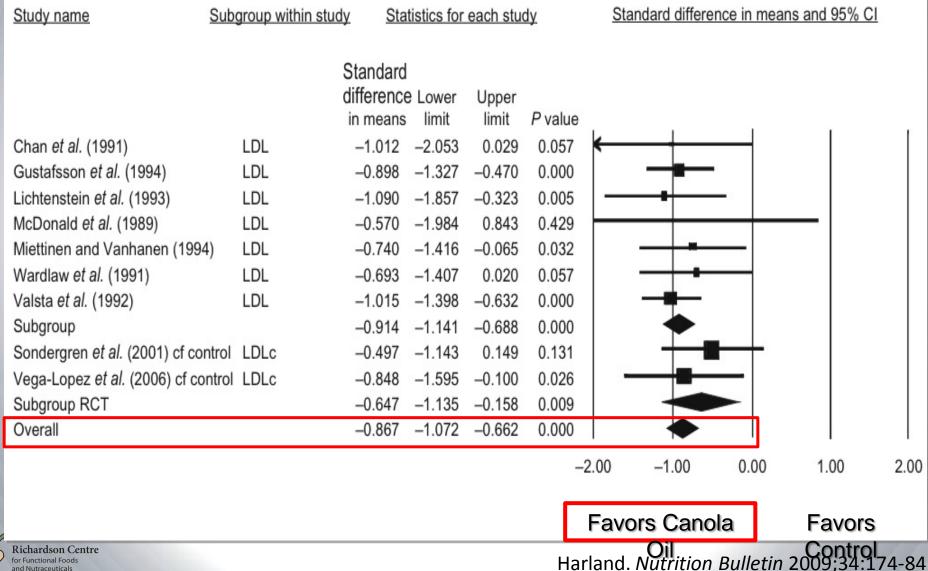
Mozaffarian & Clarke, Eur J Clin Nutr 2009;63:S22-S33

#### Mean Difference in Total-Cholesterol in Canola Oil Intervention Studies

Study name	Subgroup within stud	y Statistics for	r each st	udy	Star	ndard differe	ence in mea	ns and 95% (	21
		Standard difference Lower in means limit	Upper limit	P value					
Chan et al. (1991)	TC	-1.256 -2.328	-0.184	0.022	<	•	- 1		- 1 1
Gustafsson et al. (1994)	TC	-0.815 -1.240	-0.390	0.000			·		
Lichtenstein et al. (1993)	TC	-1.016 -1.777	-0.256	0.009		-	-		- 11
McDonald et al. (1989)	TC	-0.796 -2.236	0.643	0.278	<u> </u>			-	- 11
Miettinen and Vanhanen (1994)	TC	-0.926 -1.613	-0.238	0.008	-		-		- 11
Pedersen et al. (2000)	TC	-1.176 -1.883	-0.468	0.001					- 11
Wardlaw et al. (1991)	TC	-0.758 -1.475	-0.040	0.038	-		—		- 11
Valsta et al. (1992)	TC	-0.928 -1.308	-0.548	0.000					- 11
Subgroup		-0.924 -1.139	-0.709	0.000		•			- 11
Sondergren et al. (2001) cf contro	I TCc	-0.189 -0.826	0.448	0.561		— —			- 11
Vega-Lopez et al. (2006) cf contro	ol TCc	-0.972 -1.729	-0.215	0.012	<u> </u>	-	-		- 11
Subgroup RCT		-0.514 -1.001	-0.026	0.039					- 11
Overall		-0.857 -1.054	-0.660	0.000					
				-	2.00	-1.00	0.00	1.00	2.00
				Г	Favo	ors Cano	ola	Favor	s
Richardson Centre for Functional Foods and Nutraceuticals				Ha	rland	<b>Oil</b> Nutrition	n Bulletir	2009;34	74-84

00

#### Mean Difference in LDL-Cholesterol in **Canola Oil Intervention Studies**



and Nutraceuticals

### USDA 2006: Qualified Health Claim for Canola Oil



U.S. Food and Drug Administration



**CENTER FOR FOOD SAFETY AND APPLIED NUTRITION** 

FDA Home Page | CFSAN Home | Search/Subject Index | Q & A | Help

CFSAN/Office of Nutritional Products, Labeling, and Dietary Supplements October 6, 2006

> Qualified Health Claims: Letter of Enforcement Discretion -Unsaturated Fatty Acids from Canola Oil and Reduced Risk of Coronary Heart Disease (Docket No. 2006Q-0091)

"Limited and not conclusive scientific evidence suggests that eating about 1½ tablespoons (19 grams) of canola oil daily may reduce the risk of coronary heart disease due to the unsaturated fat content in canola oil. To achieve this possible benefit, canola oil is to replace a similar amount of saturated fat and not increase the total number of calories you eat in a day."



http://www.fda.gov/Food/LabelingNutrition/LabelClaims/QualifiedHealthClaims/ucm072958.htm

## **Composition of Canola Oil**

Canola Oil

High in omega-3 fat

**High in monounsaturated fat** 

High in plant sterols

Antioxidant vitamin E

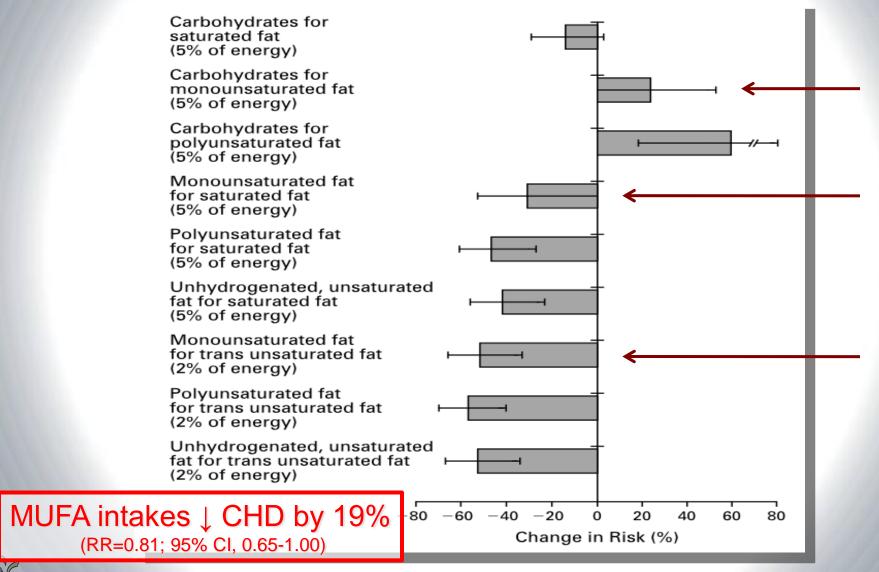
Low in saturated fat

• Other health-promoting constituents



#### **Nurses' Health Study**

14 year follow-up, n=80,082 women Estimated % change in CHD risk with dietary 5% isocaloric substitution



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Hu et al., N Engl J Med 1997;337:1491-9

## **MUFAs & CHD Risk Reduction**

Table 5. Summary of the Evidence of a Causal Association Between Diet and Coronary Heart Disease, as Determined From Examination of Prospective Cohort Studies Using the Bradford Hill Guidelines and Consistency With Findings From RCTs<sup>a</sup>

Evidence of a Causal Association			Andrew Mente, PhD; Lawrence
From Cohort Studies	Cohort Data Only	Supported by RCTs	
Strong "Mediterranean" diet <sup>b</sup>		Yes	Background: Although a w etary factors and coronary heart of the evidence supporting val
High-quality diet	-		evaluated systematically in a
Vegetables			Methods: We conducted
Nuts			MEDLINE for prospective o tzed trials investigating dieta
Trans–fatty acids			CHD. We used the Bradford
Glycemic index or load			causation score based on 4 tency, temporality, and cohe
"Prudent" diet <sup>c,d</sup>			posure in cohort studies and
"Western" diet <sup>d,e</sup>			with the findings of random
Monounsaturated fatty acids <sup>d</sup>	~		Results: Strong evidence supp
Moderate			teria satisfied) of protective fact
Fish		No	etables, nuts, and "Mediterrane patterns with CHD, and associ
Marine ω-3 fatty acids		Yes	cluding intake of trans-faity ac
Dietary folate			centic index or load. Among str
Supplementary folate		RCT data only	-22
Whole grains	200		
Dietary vitamin E			
Dietary beta carotene			
Supplementary beta carotene		RCT data only	
Dietary vitamin C			
Alcohol, light/moderate consumption			
Alcohol, heavy consumption			
Fruits			
Fiber			
Weak			
Supplementary vitamin E		Yes	
Supplementary ascorbic acid		Yes	
Total fat		Yes	
Saturated fatty acids			
Polyunsaturated fatty acids		Yes	
ω-3 Fatty acids, total		Nof	
Meat			
Eggs	~		
Milk	IIEA Statel		
	UFA intak		D eveni
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		0 00. 000/ 6	

**REVIEW ARTICLE** 

A Systematic Review of the Evidence Supporting a Causal Link Between Dietary Factors and Coronary Heart Disease

); Lawrence de Koning, MSc; Harry S. Shannon, PhD; Sonia S. Anand, MD, PhD, FRCPC

Prospective cohort studies rough a we & RCTs were used to evaluate corting vali cally in a si dietary exposures related to pective co ting dietar Bradford sed on 4 c CHD and coher udies and f randomiz ▶223 prospective studies; 66 tective facto editerranea and associa RCTs; 305 other study ts-faity acid Among stud designs Bradford Hill Guidelines score based on 4 criteria:

✓ Strength

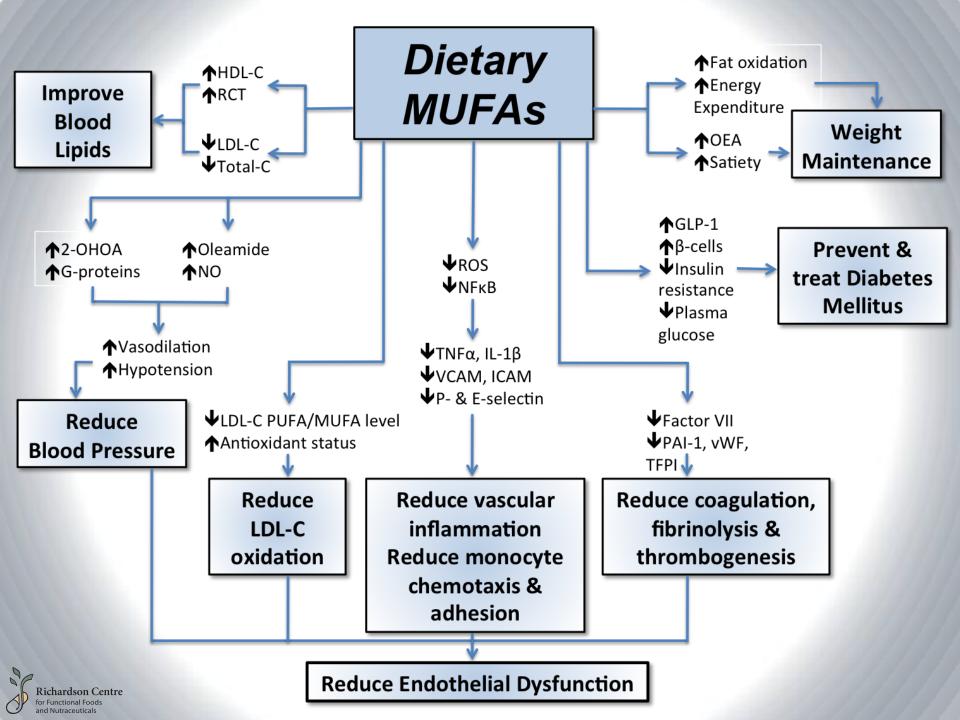
 $\checkmark$  consistency

✓ temporality

#### ents by 20% ce (RR=0.80; 95% CI, 0.67-0.93)

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Mente et al., Arch Intern Med 2009;169:659-69



## **MUFAs Improves CVD Risk Factors**

#### Blood Lipids

Harland. Nutrition Bulletin 2009; Mensink et al. Am J Clin Nutr 2003

#### Blood Pressure

Swain et al. J Am Diet Assoc 2008; Hall. Nutr Rev 2009

#### Insulin Sensitivity

Uusitupa et al. Am J Clin Nutr 1994; Tierney & Roche. Mol Nutr Food Res 2007

#### Weight Management and Body Composition

Bergouignan et al. Prog Lipid Res 2009

#### Endothelial Dysfunction

Perez-Jimenez et al. Atherosclerosis 2002

#### - LDL Oxidation Susceptibility

Egert et al. Eur J Clin Nutr 2007; Nielsen Br J Nutr 2002

#### Platelet aggregation

Smith et al. Br J Nutr 2009

References: Canola Oil Studies (Red) MUFA Review Articles (Black)

#### MUFA rich diet; hypertension, lipids, and estimated CHD risk

Characteristics of the Diet Patterns Tested in the Optimal Macronutrient Intake Trial to Prevent Heart Disease (OmniHeart): Options for a Heart-Healthy Diet

JANIS F. SWAIN, MS, RD; PHYLLIS B. McCARRON, MS, RD; EILEEN F. HAMILTON, DTR; FRANK M. SACKS, MD; LAWRENCE J. APPEL, MD

Table 3. The Optimal Macronutrient Intake Trial to Prevent Heart Disease (OmniHeart) risk factor measures and mean changes from bore (11) by diet pattern<sup>a</sup>

Clinical measure/risk	n	Baseline	CARB <sup>b</sup>	PROT <sup>c</sup>	UNSAT <sup>d</sup> <
		mean±SD°	←m	ean change from baseline±s	SD→
Systolic Blood Pressure					
(mm Hg)					
All	164	131.2±9.4	-8.2±-9.6 to -6.8	-9.5±-10.9 to -8.2	-9.3±-10.6 to -8.0
Hypertension, stage 1 <sup>f</sup>	32	146.5±5.7	-12.9±-16.6 to -9.2	-16.1±-19.7 to -12.5	-15.8±-19.4 to -12.3
LDL <sup>g</sup> cholesterol (mg/dL) <sup>h</sup>					
All	161	$129.2 \pm 32.4$	-11.6±-14.6 to -8.6	-14.2±-17.5 to -10.9	-13.1±-16.4 to -9.8
≥130 (mg/dL) <sup>h</sup>	32	156.7±21.0	-19.8±-24.2 to -15.5	-23.6±-28.5 to -18.8	-21.9±-26.9 to -16.8
HDL <sup>i</sup> cholesterol (mg/dL) <sup>h</sup>	164	50.0±16.1	-1.4±-2.5 to -0.3	-2.6±-3.6 to -1.6	-0.3±-1.3 to 0.7
Triglyceride (mg/dL) <sup>1</sup>	164	101.5±75 to 159	0.1±−8.6 to 8.8	-16.4±-25.5 to -7.3	-9.3±-17.5 to -1.2
Estimated 10-y coronary					
heart disease risk <sup>k</sup> (%)		5.1	4.3	4.0	4.1

"Moderate replacement of carbohydrate with either protein or unsaturated fat further reduced CVD risk"

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Swain et al., *J Am Diet Assoc* 2008;108:257-65

Canola

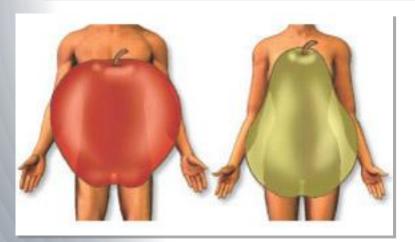
and olive

## **MUFAs & Body Fat Distribution**

Clinical Care/Education/Nutrition

#### Monounsaturated Fat-Rich Diet Prevents Central Body Fat Distribution and Decreases Postprandial Adiponectin Expression Induced by a Carbohydrate-Rich Diet in Insulin-Resistant Subjects

J.A. PANIAGUA, MD, PHD<sup>1,2</sup> A. GALLEGO DE LA SACRISTANA, MD<sup>1</sup> I. ROMERO, PHD<sup>1</sup> A. VIDAL-PUIG, MD, PHD<sup>3</sup> J.M. LATRE, MD, PHD<sup>4</sup> E. SANCHEZ, MD<sup>1</sup> P. PEREZ-MARTINEZ, MD, PHD<sup>1,2</sup> J. LOPEZ-MIRANDA, MD, PHD<sup>1,2</sup> F. PEREZ-JIMENEZ, MD, PHD<sup>1</sup> positive energy balance, which leads to obesity, is associated with insulin resistance and an increased risk of type 2 diabetes. According to our studies in rodents, adipose tissue expandability seems to be a key



**Results** – After the CHO-rich diet, subjects fat mass was redistributed from the periphery to the abdomen, compared to the MUFA-rich diet. Furthermore, the CHO-rich diet decreased postprandial adiponectin levels and insulin sensitivity, compared to the MUFA-rich diet.

Paniagua et al., Diabetes Care 2007;30:1717-23



## **Composition of Canola Oil**

High in monounsaturated fat **Canola Oil** High omega-3 fat High in plant sterols Antioxidant vitamin E Low in saturated fat

Other health-promoting constituents



## Fatty Acid Composition of Canola Oil

#### **Nutrition Facts**

Amount per serving	<b>Canada</b> 10 mL (2 tsp.)	<b>USA</b> 14 g (1 Tbsp.)
Calories	80	120
Fat	9 g	14 g
Saturated + trans	0.6 g 0 g	1 g 0 g
Polyunsaturated Omega-6 Omega-3	3 g 2 g 0.9 g	4 g 2.8 g 1.3 g
Monounsaturated	6 g	8 g
Cholesterol	0 g	0 g

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#### Prospective Cohort Studies of ALA Intakes and CHD Risk

Study	Subjects	Intakes of ALA	RR (95% CI or <i>P</i> value)
MRFIT Dolecek et al., 1992	6,250 men	Quintile extremes	↓ 40% ( <i>P</i> < 0.04)
Health Professionals Ascherio et al., 1996	43,757 men	个 1% Energy	↓ 59% (20-79%)
Finland ATBC Pietinen et al., 1997	21,930 male smokers	Quintile extremes	No significant association
Nurses Health Hu et al., 1999	76,283 women	1.36 vs. 0.71 g/d	↓ 45% (6-68%)
Zutphen Elderly Study Oomen et al., 2001	677 older men	Tertile extremes	No significant association
Iowa Women's Health Folsom et al., 2004	41,836 women	1.21 vs. 0.96 g/d	↓ 15% ( <i>P</i> trend 0.01)
Health Professionals Mozaffarian et al., 2005	45,722 men	↑ 1 g/d	$\downarrow$ 16% (0-29%) $\downarrow$ 47% (17-66%) for low seafood eaters

A study involving eleven eastern European countries showed that increase in ALA consumption from **rapeseed oil** attributed in reducing the CHD mortality (r = -0.84 in men and -0.83 in women) (Zatonski et al., Eur J Epidemiol 2008;23:3–10)

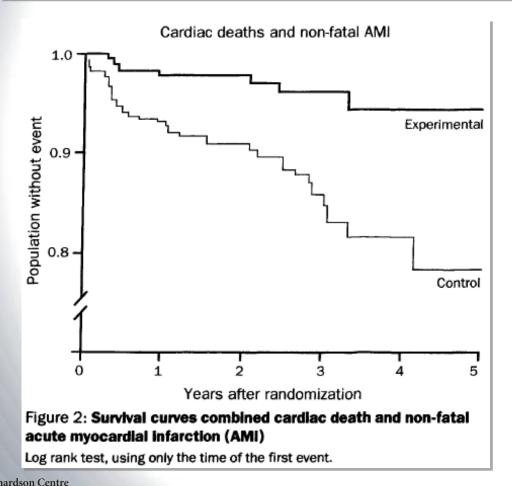


Table adapted from: Mozaffarian et al., Altern Ther Health Med 2005;11:24-30

r Functional Foods

#### Mediterranean alpha-linolenic acid-rich diet in secondary prevention of coronary heart disease

Michel de Lorgeril, Serge Renaud, Nicole Mamelle, Patricia Salen, Jean-Louis Martin, Isabelle Monjaud, Jeannine Guidollet, Paul Touboul, Jacques Delaye



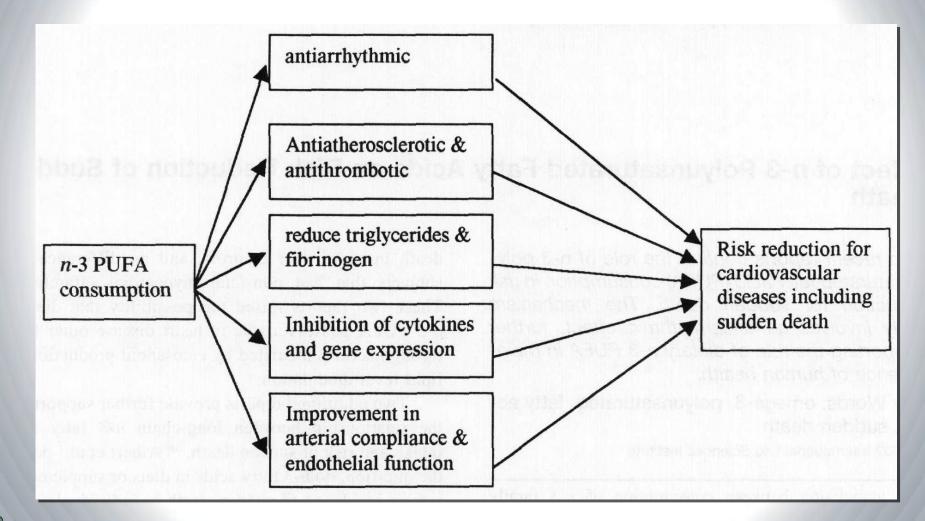
Experimental group diet added canola oil and canola oil-based margarine...

...contributing to the **3-fold increase in daily ALA intakes** Control=0.27% energy from ALA Experimental=0.81% energy from ALA

#### ↓ 73% (95% CI: 41-88%) CHD risk reduction with ALA-rich diet

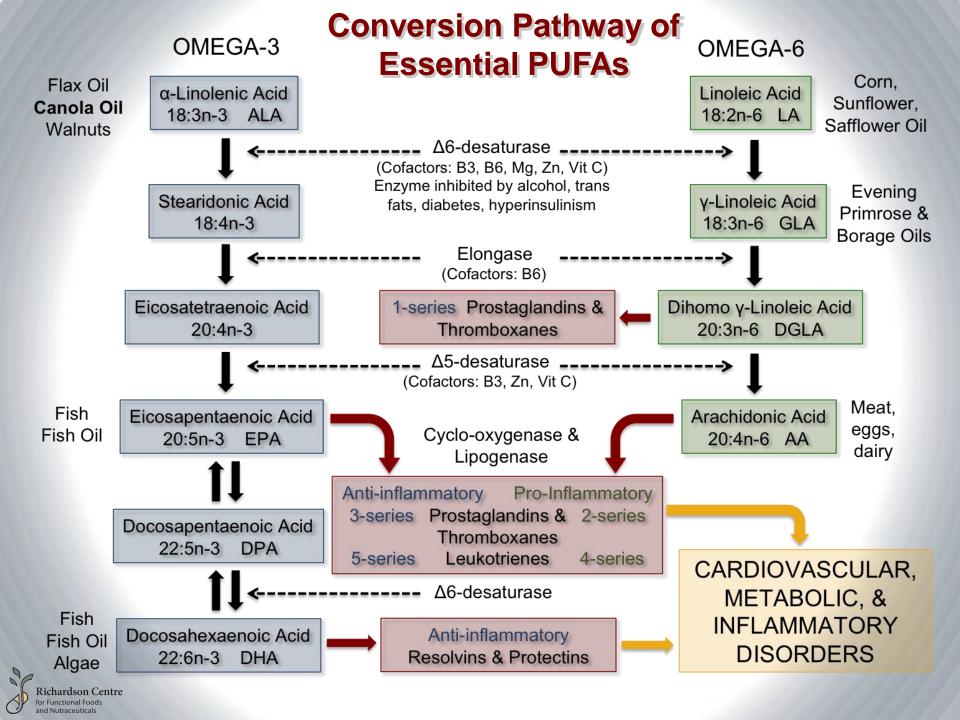
De Lorgeril et al., Lancet 1994;343:1454-59

#### Mechanisms of Action by which Omega-3 Fatty Acids may Decrease CVD Risk

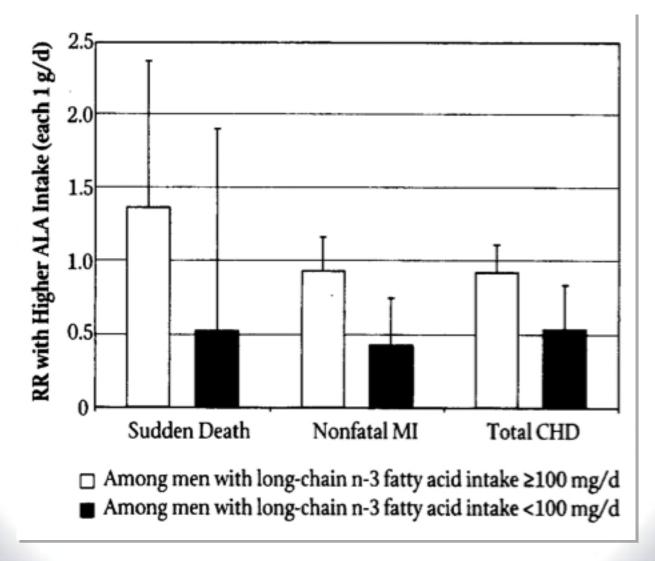


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Jones & Lau, Nutr Rev 2002;60:407-9



#### Relative Risk of CHD Associated with Each 1 g/day of High ALA Intake





Mozaffarian et al., Altern Ther Health Med 2005;11:24-30

# Alpha Linolenic Acid & Endothelial Dysfunction

	Measurements	Studies	Results
Inflammation	Arachidonic Acid (AA), PGI <sub>2,</sub> IL-6, IL-1β, TNF-α, CRP, VCAM-1, ICAM-1,	Sekine et al., 2007	$\downarrow$ vascular AA levels, $\uparrow$ PGI <sub>2</sub> formation
	E-Selectin, fibrinogen, Serum Amyloid A (SAA)	Zhao et al., 2007	$\downarrow$ IL-6, IL-1 $\beta$ , and TNF- $\alpha$ production
		Wendland et al., 2006	$\downarrow$ fibrinogen levels
		Lopez-Garcia et al., 2004	$\downarrow$ CRP, IL-6, E-Selectin
		Zhao et al., 2004	↓ CRP, VCAM-1, ICAM- 1, E-Selectin
		Rallidis et al., 2004	$\downarrow$ VCAM-1
		Bemelmans et al., 2004	↓ CRP
		Rallidis et al., 2003	↓ CRP, SAA, IL-6

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## **Composition of Canola Oil**

High in monounsaturated fat **Canola Oil** High in omega-3 fat High in plant sterols Antioxidant vitamin E Low in saturated fat

Other health-promoting constituents



## **Plant Sterols in the Diet**

#### Average daily plant sterol intake of adults 150 - 400 mg/day

#### major source:

- fat and oils (~1g/100ml canola oil)
- bread and cereals
- fruits and vegetables
- nuts

Recommended intake of plant sterol-enriched foods for a significant cholesterol-lowering effect **2 g/day** 







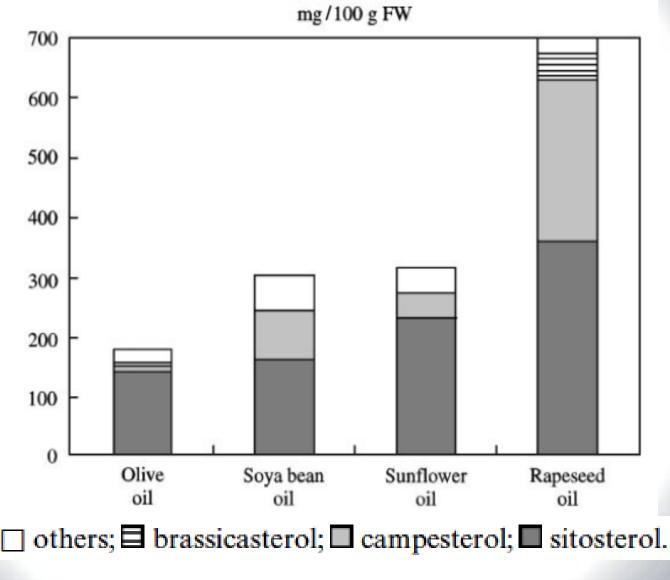






Katan et al., *Mayo Clin Proc* 2003;78:965-78; National Cholesterol Education Program (NCEP) Expert Panel JAMA 2001 May 16;285:2486-97

#### **Plant Sterol Content of Refined Vegetable Oils**

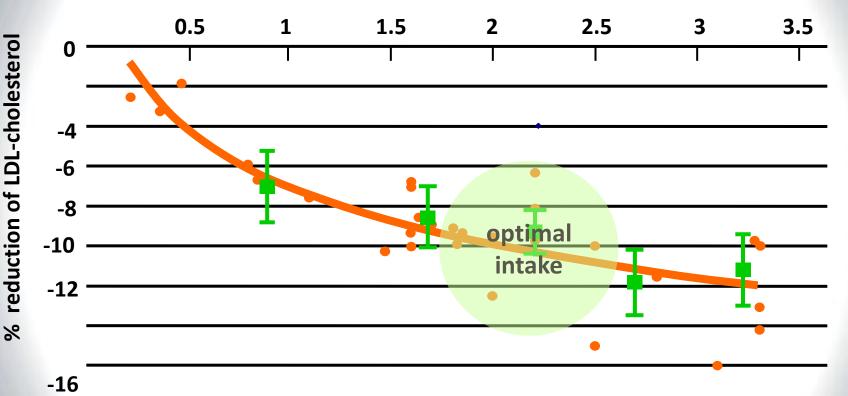


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Piironen et al., JFCA 2000;13:619-24

#### Cholesterol Lowering with Plant Sterols in Fat-based Foods: Dose-response Relationship

Plant sterol intake (g/day)



data of ~ 30 placebo-controlled Unilever initiated studies with phytosterol-enriched spreads

data (mean plus 95% confidence interval) from meta-analysis of 41 studies with phytosterols or stanols

ichardson Centre

for Functional Foods and Nutraceuticals Adapted from: Katan et al., Mayo Clin Proc. 2003;78:965-978

An olive oil-rich diet results in higher concentrations of LDL cholesterol and a higher number of LDL subfraction particles than rapeseed oil and sunflower oil diets

> Anette Pedersen,<sup>1,\*</sup> Manfred W. Baumstark,<sup>†</sup> Peter Marckmann,<sup>\*</sup> Helena Gylling,<sup>§</sup> and Brittmarie Sandström<sup>\*</sup>

**Design:** Randomized crossover trial; 18 healthy men; 3 week dietary intervention **Diet:** Olive oil, rapeseed oil (canola oil), or sunflower oil (50 g oil per 2500 kcal/d)

**Results:** As compared to the olive oil and sunflower oil diets, the rapeseed oil (canola oil) diet resulted in the most favorable effects on:

- Plasma lipids, including LDL and VLDL
- Lipids ratios, including TC:HDL and LDL:HDL
- Plasma apolipoproteins
- Number and lipid content of LDL subfractions, ie VLDL

#### "...Some of the differences may be attributed to differences in the squalene and phytosterol contents of the



JOURNAL OF LIPID RESEARCH

Pedersen et al., J Lipid Res 2000;41:1901-11

http://www.stockton-press.co.uk/ejcn

#### Review

# Monounsaturated oils do not all have the same effect on plasma cholesterol

AS Truswell and N Choudhury

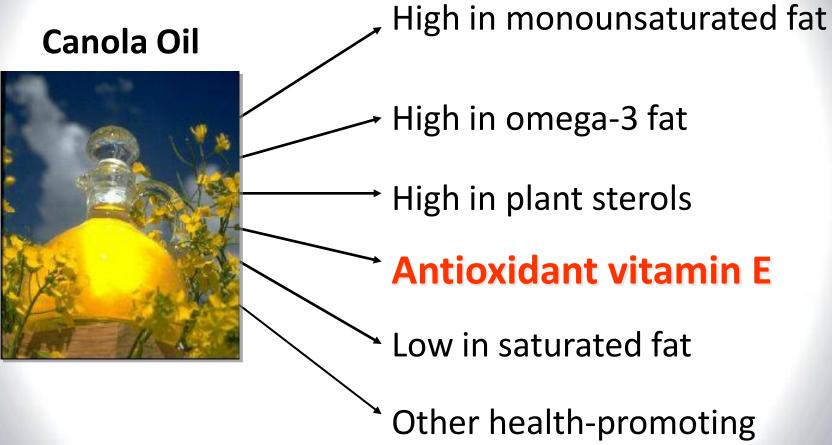
Human Nutrition Unit, University of Sydney, Sydney, NSW 2006, Australia

...it has been found that olive oil has little to no effect on cholesterol lowering, as compared to other MUFA oils, such as **canola oil** or high-oleic sunflower oil



Truswell & Choudhury. Eur J Clin Nutr 1998;52:312-15

## **Composition of Canola Oil**



constituents



## **Canola Oil Contains Vitamin E**





#### 1 Tbsp. of Canola Oil provides ~2.9 mg of Vitamin E

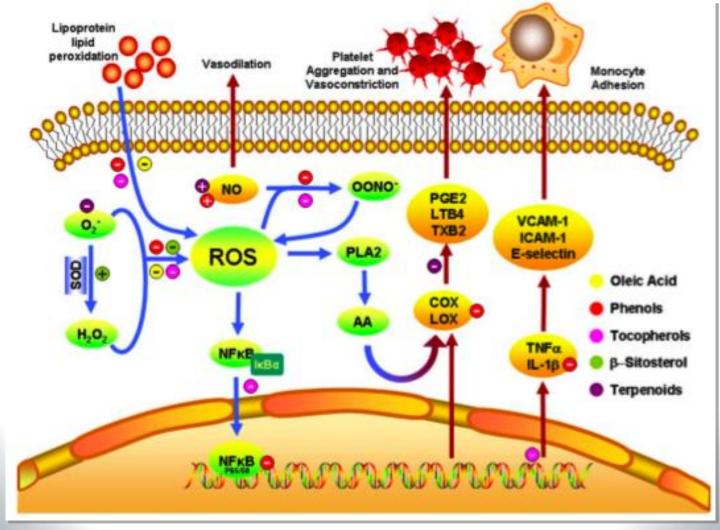
➤ equivalent to ~1/5 of the recommended daily intake for adults (15 mg ATE\*).

Vitamin E present in canola oil could be beneficial in the prevention and treatment of diseases related to oxidative stress including cancer, cardiovascular and neurodegenerative disorders



Giugliano. Nutr Metab Cardiovasc Dis 2000;10:38-44; Practico & Delanty. Am J Med 2000;109:577-85

### Endothelial Function: Proposed Mechanisms of Action of Oleic Acid and Other Minor Compounds from Vegetable Oils



Richardson Centre for Functional Foods and Nutraceuticals

Perona et al., J Nutr Biochem 2006;17:429-45



### **Richardson Centre** for Functional Foods and Nutraceuticals



Canola and flax oils in modulation of plasma lipids, vascular function and biomarkers of cardiovascular disease risk; Randomized crossover controlled trials

#### Canola & Flax; Clinical Trial 1

- Sept 2007 2010
- Richardson Centre (RCFFN) at Univ. of Manitoba
- 3 treatments (4 weeks)
  - High-oleic Canola Oil
  - HOCO/Flaxseed Oil blend
  - Average American Diet (control)
- 36 hyperlipidemic subjects
- Clinical trial complete
- Analysis in progress

#### Canola & Flax; Clinical Trial 2

- Sept 2010 2012
- Multi-Centre Trial
  - RCFFN (Univ. of Manitoba),
     Univ. of Toronto, Laval Univ., &
     Penn State Univ.
- 5 treatments (4 weeks)
  - Canola oil
  - DHA enriched canola oil
  - High-oleic acid canola oil
  - Flax oil
  - Corn oil (control)
- 140 subjects (35 per site)



# **Clinical Trial Endpoint Analysis**

#### Endothelial health and body composition

- Flow-mediated dilation (by Endo-PAT2000)
- Body fat deposition by DEXA
- Plasma lipids and lipoproteins, inflammatory, & peroxidation biomarkers
  - TC, HDL-C, LDL-C, and TG, lipoprotein subclasses
  - CRP, IL-6, IL-10, sTNFRα, sVCAM-1, sICAM-1, E-selectin
  - Urinary isoprostanes and prostaglandins

#### ALA conversion to long chain n-3 fatty acids

- [U-13C] alpha-linolenic acid (70mg dose)
- FADS1/FADS2 mRNA and protein expression
- Genetic analyses of FADS1 and FADS2



## Summary

- Canola oil contains several constituents that reduce risk of cardiovascular disease and other chronic disorders
- The monounsaturated fat content of canola is associated with favorable modulation of lipid levels, blood pressure, insulin sensitivity, as well as oxidative and inflammatory status
- The omega-3 content of canola is predicted to exert desirable changes in cardiovascular risk, associated with beneficial effects on endothelial function and numerous other health-related parameters





- Plant sterols in canola oil further contribute to the reduction in LDL-C levels
- Additional bioactives in canola oil may exert positive biological effects on health, such as vitamin E
- Clinical studies on canola oil are currently being conducted to substantiate the cardioprotective benefits of canola oil







#### **Richardson Centre**

for Functional Foods and Nutraceuticals

## Thank you





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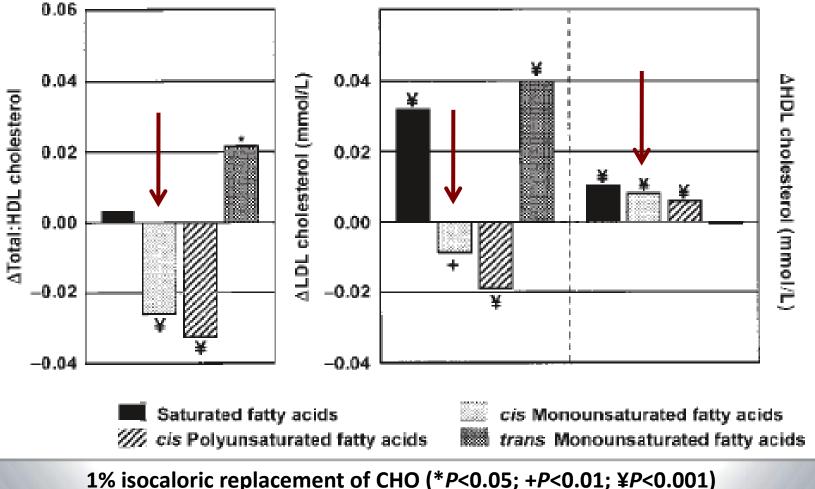
### **APPENDIX**

### **Supplemental Material**



### **MUFAs & Blood Lipids**

Effects of dietary fatty acids on serum lipids: A meta-analysis of 60 controlled trials

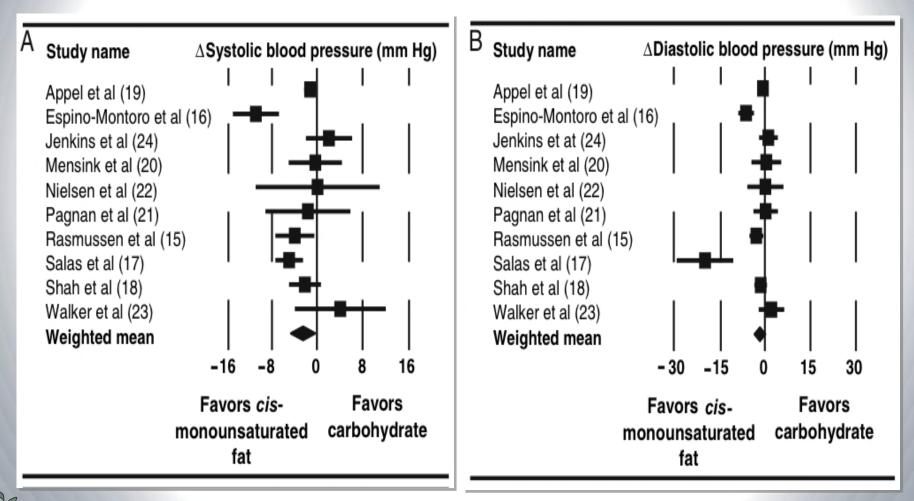


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for Functional Foods and Nutraceuticals Mensink RP et al., Am J Clin Nutr 2003;77:1146-55

## **MUFAs & Blood Pressure**

#### Effect of high-CHO or high-*cis*-MUFA fat diets on blood pressure: A meta-analysis of interventions trials



Richardson Centre

and Nutraceuticals

Shah et al., Am J Clin Nutr 2007;85:1251-6

### **Effects of MUFA on LDL Oxidative Status**

Reference	Subject	Design/duration	Diets	Outcome (MUFA vs. other diets)
Moreno et al. 2008	Healthy men (n=20)	Randomized,CO 4 weeks	MUFA SFA H-CHO	<ul> <li>↑ OxLDL lag time</li> <li>↑ propagation rate</li> <li>↓ oxLDL macrophage uptake</li> </ul>
Ahuja et al. 2003	Healthy sbj (n=31)	Randomized, CO 14-16 days	MUFA H-CHO	个 OxLDL lag time
Nielsen et al. 2002	Healthy sbj (n=18)	Randomized, CO 3 weeks	MUFA n-3 PUFA* n-6 PUFA	<ul> <li>↑ OxLDL lag time</li> <li>↓ propagation rate with MUFA</li> <li>and n-3 PUFA vs. PUFA</li> </ul>
Hargrove et al. 2001	Healthy sbj (n=20)	Randomized, CO 3.5 weeks	AAD NCEP 3 MUFA diets	<ul> <li>↑ OxLDL lag time</li> <li>↓ rate of OxLDL</li> </ul>
Ashton et al. 2001	Healthy sbj (n=28)	Randomized, CO 1 month	MUFA H-CHO	<ul> <li>↑ OxLDL lag time</li> <li>↓ rate of oxidation</li> <li>↓ conjugated dienes</li> </ul>
Baroni et al. 1999	HC patients (n=13)	CO 8 weeks	MUFA PUFA	个 OxLDL lag time

### **MUFA & Endothelial Dysfunction**

	Measurements	Studies	Results
Inflammation & Hemostasis	Acute Phase Reactants: • CRP, Fibrinogen	Mena et al., 2009	↓ IL-6, ICAM-I, VCAM-I & CRP
	<b>Cytokines:</b> • IL-6, TNF-α	Pacheco et al., 2008	↓ post-prandial AUC for ICAM-I & VCAM-I
	Adhesion Molecules:	Bogani et al., 2007	$\downarrow$ TXB <sub>2</sub> & LTB <sub>4</sub> , $\uparrow$ PAC
	• VCAM-I, ICAM-I, E- selectin	Brunelleschi et al., 2007	$\downarrow$ NF-κB translocation
	Coagulation and	Pacheco et al., 2006	↓tissue factor, fibrinogen, PAI-1
	Fibrinolysis Factors: • Tissue factor, Factor	Serrano-Martinez et al., 2005	$\downarrow$ TNF-α & VCAM-I
	VIIc, TXB <sub>2</sub> , LTB <sub>4</sub> , PAI-1, Platelet aggregation	Visioli et al., 2005	$\downarrow$ TXB <sub>2</sub> & $\uparrow$ PAC
	Others:	Allman-Farinelli et al., 2005	$\downarrow$ factor VIIc
	• NF-κB, Plasma antioxidant capacity	Smith et al., 2003*	$\downarrow$ platelet aggregation & $\downarrow$ factor VIIc
0	(PAC)	Kwon et al., 1991*	$\downarrow$ AA & platelet aggregation

### **MUFA Effects on Insulin & Glucose Responses**

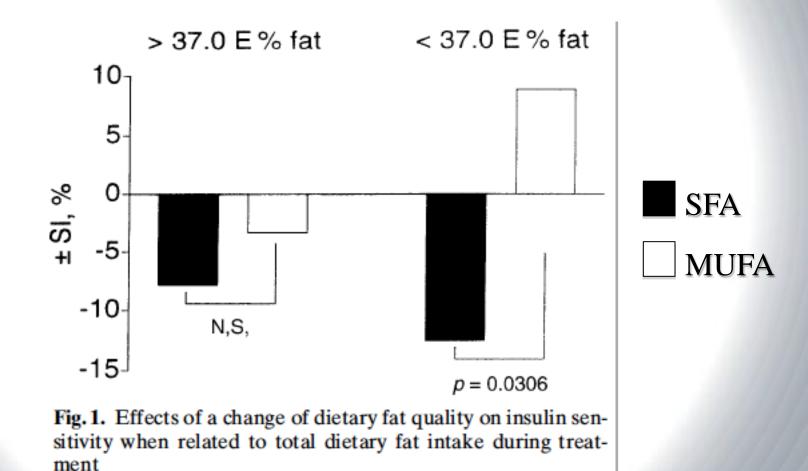
Reference	Subject (sbj)	Design/duration	Diets	Outcome (MUFA vs. other diets)
Due et al. 2008	Obese sbj (n=46)	Randomized, PAR 6 months	MUFA SFA Low-fat	<ul> <li>↓ HOMA-IR</li> <li>↓ fasting glucose, insulin</li> </ul>
Lopez et al. 2008	Healthy men (n=14)	Randomized, CO Single meal	4 diets varying in MUFA:SFA	个 postprandial β-cell function and insulin sensitivity with 个 in MUFA:SFA of diet
Shah et al. 2007	T2DM Subjects (n=11)	Randomized, CO Single meal	SFA MUFA n-6 PUFA EPA+DHA	<ul> <li>↓ postprandial insulin response vs.</li> <li>SFA and n-6 PUFA</li> <li>↔ postprandial glucose response</li> </ul>
Paniagua et al. 2007	Obese T2DM (n=11)	Randomized, CO 28 days	SFA MUFA H-CHO	<ul> <li>↑ insulin sensitivity</li> <li>↓ fasting glucose</li> <li>↑ postprandial GLP-1</li> </ul>
Vessby et al. 2001	Healthy sbj (n=162)	Randomized 3 months	SFA MUFA	个 insulin sensitivity
Joannic et al. 1997	Healthy sbj (n=8)	Randomized, CO Single meal	<b>MUFA*</b> n-6 PUFA	个postprandial glucose and insulin responses

Abbreviations: CO, crossover; PAR, parallel arm; T2DM Type 2 Diabetes Mellitus; GLP-1, glucagon-like peptide-1



## **MUFAs & Insulin Sensitivity**

# Substituting dietary SFA for MUFA impairs insulin sensitivity in health men and women: The KANWU study

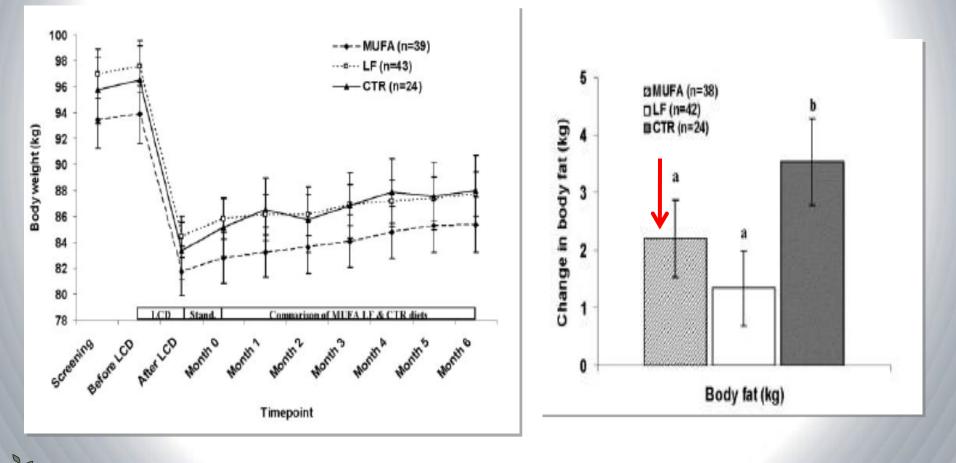


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Vessby et al., Diabetologia 2001;44:312-19

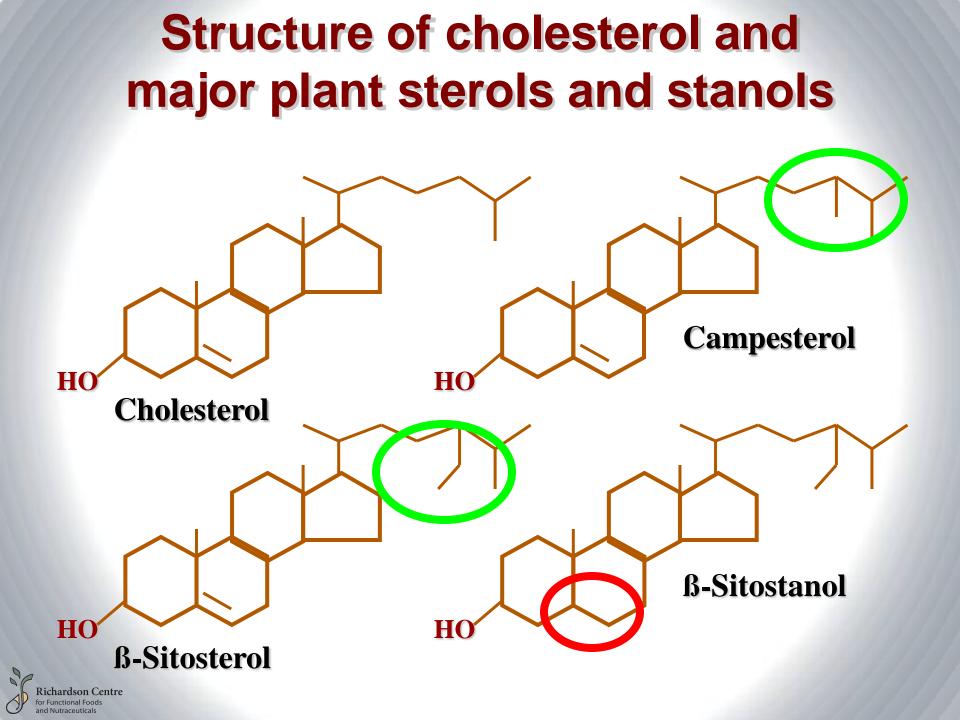
## **MUFAs & Weight Management**

#### Change in Body Weight and Body Fat at Baseline and 6 Months: Effects of MUFA, Low Fat, & Control Fat Diets



Richardson Centre for Functional Foods and Nutraceuticals

Due et al., Am J Clin Nutr 2008;88:1232-41



### Cholesterol-lowering effect of plant sterols: mechanism of action

Intake of 2 g plant sterols/day  $\rightarrow$  30-40% reduction in cholesterol absorption

