

Olive oil consumption and risk of breast cancer in the Canary Islands: a population-based case–control study

Purificación García-Segovia^{1,2}, Almudena Sánchez-Villegas¹, Jorge Doreste¹, Francisco Santana¹ and Lluís Serra-Majem^{1,3,*}

¹Department of Clinical Science, University of Las Palmas of Gran Canaria, Apdo. Correos 550, 35080 Las Palmas G/C, Spain: ²Department of Food Technology, Polytechnical University of Valencia, Spain: ³Foundation for the Advancement of Mediterranean Diet, University of Barcelona Science Parc, Spain

Abstract

Background: Breast cancer mortality and incidence rates in the Canary Islands, and particularly in Gran Canaria, are higher than those in the rest of Spain.

Objectives and design: A case–control study was designed to assess the role of differential fatty acid intakes and olive oil consumption on breast cancer risk in the Canary Islands. The study was conducted between 1999 and 2001, including a total of 755 women: 291 incident cases with confirmed breast cancer and 464 controls randomly selected from the Canary Island Nutrition Survey (ENCA). A semi-quantitative food-frequency questionnaire was completed and potential confounders were adjusted using unconditional logistic regression.

Results: Compared to the first quintile of intake, the highest quintile of monounsaturated fat intake was significantly related to a lower risk of breast cancer (odds ratio = 0.52; 95% CI 0.30–0.92). Regarding olive oil, the odds ratio for women in the three upper quintiles of consumption (≥ 8.8 g/day) was 0.27 (95% CI 0.17–0.42).

Conclusion: Our results support the protective role of olive oil consumption on breast cancer among Canaries women.

Keywords
Breast cancer
Olive oil
Fat intake
Case–control study

Evidence on dietary fat intake and breast cancer risk come from animal experimentation, ecological and migrant studies, analyses of secular trends and analytical designs such as case–control and cohort studies¹. Whereas animal studies and international comparisons have suggested a positive association between total fat intake and the risk of breast cancer, particularly among postmenopausal women, longitudinal studies have failed to confirm the deleterious effect of overall lipid intake or specific types of fatty acids on the disease^{2–4}.

Ecological studies carried out in Mediterranean regions have reported a lower incidence of breast cancer as compared with those from Northern European countries or North America⁵. In spite of the high total fat intake observed in the Mediterranean area, around 30 or 40% of total energy intake, this paradoxical finding has been explained by its higher monounsaturated/saturated fat ratio⁶. Moreover, several studies performed in Southern Europe have suggested that olive oil, which is the main source of fat used to dress salads and to cook, could exert a protective role on breast cancer^{7–10}.

It has been estimated that countries like Greece and Spain obtain approximately 25 and 22% of total energy, respectively, from vegetable oil consumption, olive oil included¹¹. Olive oil is one of the most important features

of the Mediterranean diet, which has been postulated as being protective against several diseases including coronary heart disease and cancer. Additionally, olive oil has been associated with lower mortality rates in these regions.

Whereas, breast cancer mortality and incidence are lower in Spain than in other European countries or in the USA, there are alarming rates in the Canary Islands and particularly in Gran Canaria. Thus, the aim of our study was to ascertain the effect of olive oil consumption and different fat intake patterns on breast cancer risk in the Canary Islands, where legislative initiatives and marketing have promoted olive oil consumption in the past 20 years.

Methods

Case and control assessment

A case–control study designed to evaluate the relationship between diet and breast cancer was conducted between April 1999 and June 2001 in Las Palmas and Santa Cruz de Tenerife. During the study period, 326 cases (women with a histological confirmed first diagnosis of breast cancer) were collected from the Canary Islands teaching hospitals. We also selected 492 controls among women aged 25–85 years from the Canary Islands Nutrition Survey (Encuesta de Nutrición de Canarias, ENCA) participants.

*Corresponding author: Email lserra@dcc.ulpgc.es

Exposure assessment

A face-to-face interview was conducted in hospital (cases) or primary care settings (controls) by trained personal. A semi-quantitative food-frequency questionnaire (SFFQ, 88 items) was used to assess the usual diet of the participants, total energy and macro- and micronutrient intakes. It was modified from a previously validated questionnaire¹² to take into account the Canarian diet features.

The food items were categorised into the following groups: cereals and pulses, dairy products, fruits, vegetables, meat, eggs and fish, olive oil and other fats, sweets, nuts, and beverages (including alcoholic drinks). The questionnaire also included data on dressing, cooking and frying fats, to describe the fat intake pattern.

The food composition database was established on the basis of the Spanish food composition tables¹³.

A lifestyle questionnaire asked for sociodemographic variables, chronic diseases, smoking habits (age at onset, age at quitting, number of cigarettes smoked per day, type of tobacco), alcohol consumption (type of beverages, amount of alcohol consumed per day, drinking patterns), physical activity, family history of diseases, menstrual and reproductive events and nutritional beliefs, opinions and attitudes.

Anthropometric variables were collected; women were weighed and measured without shoes and outer clothing.

Statistical analysis

Women with missing values for the main variables or out of *a priori*-defined daily energy intake range¹⁴ ($< 1.2 \times$ estimated resting metabolic rate – ≥ 4000 kcal/day) were excluded from the analysis. The final study population was composed of 755 women (cases = 291, controls = 464).

The nutrient residual model approach¹⁵ was used to control for the potential confounding effect of energy intake on olive oil consumption and fatty acids intake.

The exposure variables were categorised into quintiles all over the entire study group distribution. The associations were measured fitting unconditional logistic regressions. The lowest quintile of intake was used as the reference category. Odds ratios (OR) and 95% confidence intervals (CI) were calculated. Models were adjusted for age (years), smoking status (never smoked, ex-smoker and smoker), education level (primary or less, secondary and university degree), presence/absence of benign breast disease, pre- or postmenopausal status, and body mass index (kg/m^2 , and a quadratic term for body mass index to account for non-linearity). Additionally, every fatty acid intake was adjusted for each other. Finally, we looked for interactions in the olive oil consumption and fat intake-breast cancer associations.

SPSS[®] version 11.0 (SPSS Inc., Chicago, IL, USA) statistical software was used in the analysis.

Results

The main characteristics of the participants are shown in Table 1. The mean age was 55.5 years for the cases and 53.1 years for the controls. There were differences according to menopausal status (postmenopausal women: cases 72.5%; controls 57.5%) as well as to benign breast disease history (cases 44.3%; controls 6.5%).

Both cases and controls reported a low level of daily physical activity. There was a statistically significant difference in smoking habits, with a higher proportion of smokers in the control group. Energy intake was higher among cases (2237.7 kcal/day) than among controls (2056.2 kcal/day).

Nutrient intakes were similar for cases and controls (Table 2). Energy intake from total fat (40.8% for cases and 40.9% for controls) and from monounsaturated fatty acids (MUFA) (16.9% and 16.7%) were almost identical.

Table 1 Characteristics of case and control women

	Cases (<i>n</i> = 291)	Controls (<i>n</i> = 464)	<i>P</i>
Age (years) [mean \pm SD]*	55.5 \pm 11.8	53.1 \pm 11.4	<0.01
Body mass index (kg/m^2) [mean \pm SD]*	29.0 \pm 5.0	28.0 \pm 5.3	0.02
Energy intake (kcal/day) [mean \pm SD]*	2237.7 \pm 616.6	2056.2 \pm 620.1	<0.001
Educational level (%)†			0.7
Primary or less	76.6	75.9	
Secondary	13.1	15.7	
University degree	10.3	8.4	
Smoking (%)†			<0.01
Never	64.9	71.3	
Ex-smoker	17.5	6.9	
Smoker	17.5	21.8	
Physical activity level (%)†			0.5
Never	74.9	71.6	
Smooth	22.0	24.1	
Moderate	3.1	4.3	
Premenopausal (%)†	28.5	42.5	<0.01
Benign breast disease (%)†	55.7	93.5	<0.01

SD, standard deviation.

*Mann–Whitney and †chi-square *P*-values.

Table 2 Mean and quintiles of total caloric and macronutrient intakes for cases and controls

	Controls (<i>n</i> = 464)					Cases (<i>n</i> = 291)				
	Mean ± SD	Percentile				Mean ± SD	Percentile			
		20th	40th	60th	80th		20th	40th	60th	80th
Energy (kcal)	2056 ± 620	1509	1844	2155	2578	2238 ± 617	1663	2059	2451	2800
Carbohydrate (g)*	211 ± 45	179	202	222	244	213 ± 47	174	203	224	256
Protein (g)*	86 ± 14	75	82	88	97	85 ± 15	75	84	90	98
Total fat (g)*	100 ± 19	85	95	103	113	97 ± 19	82	93	102	113
MUFA (g)*	41 ± 9	33	39	42	48	40 ± 9	33	38	42	47
PUFA (g)*	13 ± 4	10	11	13	15	13 ± 4	10	11	13	15
SFA (g)*	37 ± 13	27	33	38	46	35 ± 12	25	31	37	43
Fibre (g)*	22 ± 8	16	20	23	28	25 ± 8	17	22	27	31
MUFA/SFA	1.3 ± 0.9	0.9	1.0	1.2	1.5	1.3 ± 1.1	0.9	1.0	1.2	1.6
SFA (%)	14.8	10.0	12.6	15.5	19.3	14.6	10.4	13.0	15.4	18.4
PUFA (%)	5.4	4.1	4.7	5.5	6.4	5.3	4.1	4.7	5.4	6.2
MUFA (%)	16.7	13.2	15.5	17.5	20.3	16.9	13.2	16.1	17.9	20.1
Total fat (%)	40.9	33.8	38.8	43.2	47.9	40.8	33.9	39.1	42.8	47.9

MUFA, monounsaturated fatty acids; PUFA, polyunsaturated fatty acids; SFA, saturated fatty acids.

* Adjusted for energy intake.

OR (95% CI) for the associations between fatty acid intakes and breast cancer risk are shown in Table 3. Breast cancer and PUFA (polyunsaturated fatty acids) and SFA (saturated fatty acids) intakes were not associated. Linear trends were not statistically significant.

For MUFA, adjusted OR were 1.04 (0.62–1.76), 0.71 (0.42–1.23), 0.80 (0.46–1.36), and 0.52 (0.30–0.92), with only the latter (5th quintile, MUFA intake >47 g/day) being statistically significant. A significant linear trend was observed ($P = 0.02$). However, when the model was additionally adjusted for the other dietary fatty acids these associations disappeared.

Olive oil consumption was analysed separately, and we also observed an inverse linear trend with breast cancer

risk ($P < 0.001$) (Fig. 1). The OR for women in the three upper quintiles (≥ 8.8 g/day) was 0.27 (95% CI 0.17–0.42). To assess its cumulative effect, daily consumption was also considered in a quantitative basis, obtaining an OR of 0.97 (95% CI 0.96–0.98) for each additional gram of olive oil consumption.

Discussion

A higher consumption of olive oil has been found to be associated with a lower risk of breast cancer. Our results also suggest a dose–response relationship between MUFA intake and breast cancer risk, but not statistically significant after adjusting for the rest of fatty acids in the diet.

Table 3 Association between different types of fats and breast cancer

Energy-adjusted fatty acids intake (quintile)	Crude OR (95% CI)	Adjusted OR (95% CI)*	<i>P</i> trend	Additionally adjusted OR (95% CI)†	<i>P</i> trend
PUFA (g/day)					
< 10	1	1		1	
10–11	1.06 (0.67–1.67)	1.18 (0.70–2.00)	0.42	1.21 (0.70–2.09)	0.49
11–13	0.89 (0.56–1.42)	0.79 (0.46–1.37)		0.83 (0.46–1.49)	
13–15	0.95 (0.60–1.50)	0.89 (0.51–1.55)		0.94 (0.51–1.74)	
≥ 15	0.87 (0.55–1.38)	0.87 (0.51–1.51)		0.88 (0.48–1.63)	
SFA (g/day)					
< 25	1	1		1	
25–31	0.78 (0.50–1.24)	0.82 (0.47–1.42)	0.18	0.82 (0.47–1.44)	0.39
31–37	0.70 (0.44–1.11)	0.70 (0.40–1.22)		0.73 (0.42–1.28)	
37–43	0.78 (0.50–1.24)	1.09 (0.64–1.86)		1.18 (0.68–2.05)	
≥ 43	0.60 (0.38–0.96)	0.60 (0.35–1.05)		0.66 (0.37–1.21)	
MUFA (g/day)					
< 33	1	1		1	
33–38	1.15 (0.72–1.82)	1.04 (0.62–1.76)	0.02	1.14 (0.66–1.96)	0.07
38–42	0.89 (0.56–1.42)	0.71 (0.42–1.23)		0.80 (0.45–1.40)	
42–47	1.06 (0.67–1.68)	0.80 (0.46–1.36)		0.93 (0.52–1.67)	
≥ 47	0.95 (0.59–1.51)	0.52 (0.30–0.92)		0.61 (0.34–1.11)	

OR, odds ratio; CI, confidence interval; PUFA, polyunsaturated fatty acids; SFA, saturated fatty acids; MUFA, monounsaturated fatty acids.

* Logistic regression, adjusted for age, smoking, body mass index (BMI) and quadratic BMI, menopausal status and benign breast disease.

† Additionally adjusted for all other variables shown in the table.

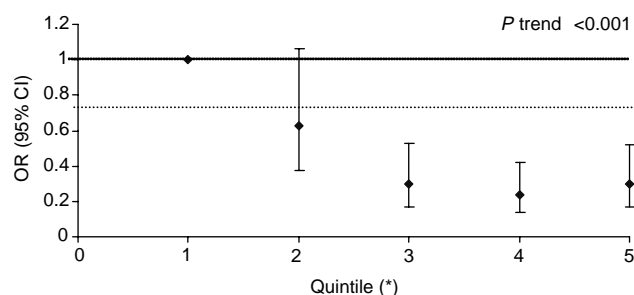


Fig. 1 Odds ratios (95% confidence intervals) for quintile of energy-adjusted olive oil consumption adjusted for age, smoking status, education level, benign breast disease, menopausal status, body mass index (BMI) and quadratic BMI. *Quintile (g/day): 1 = ≤ 3.2 ; 2 = 3.3–8.7; 3 = 8.8–16.0; 4 = 16.1–27.4; 5 = > 27.4

Epidemiological studies have frequently failed to find a relevant association between total or different type of fat intakes and risk of breast cancer^{2–4}. It has been argued that an extremely low-fat/high-carbohydrates diet is causally related to breast cancer¹⁶.

However, several studies support a protective role for MUFA intake^{17,18} and olive oil consumption^{7–10} on breast cancer risk. The results have been linked to the theory that monounsaturated olive oil's fatty acids structure hinders free radical-initiated peroxidation.

For example, in a Greek case–control study the OR for the highest consumption of olive oil relative to the 'once-a-day' reference category was 0.75 (CI 95% 0.57–0.98)⁸. Similar results were observed in a case–control study carried out in Italy (OR = 0.89 per any increment of 30 g in olive oil consumption)¹⁰.

In a multi-centre case–control study in five regions of Spain, total fat and MUFA intakes and olive oil consumption⁷ were evaluated, with only the latter being inversely associated with breast cancer (OR = 0.66, 95% CI 0.46–0.97 for the highest quartile of consumption).

EURAMIC, a multi-centre case–control study carried out in 11 European countries, reported a strong and protective effect of oleic acid intake, the most frequent monounsaturated fatty acid contained in olive oil, but only in Spain¹⁸. Consequently, it has been postulated that oleic acid intake is a proxy for other olive oil components.

Specifically, current interest is centered on polyphenolic compounds of extra-virgin olive oil (hydroxytyrosol, oleuropein, flavonoids, and catechins) that have demonstrated antioxidant properties in laboratory experiments^{19,20}.

Breast and other cancer mortality rates have incremented in the Southern European countries in the last years. These changes correlate with an increase in the consumption of milk, meat and animal fat products. In Spain, it has been accompanied with a decrease in olive oil and an increase of other vegetables oils consumption²¹.

Breast cancer incidence and mortality rates are even higher in the Canary Islands. Whereas the incidence is around 48 cases per 10⁵ person-years in Spain, the value increases to 53 cases in Las Palmas, one of the two

provinces of the Archipelago. However, the gap between the Canary Islands and the rest of Spain has been shortened in the last years. A possible explanation could be the increment in olive oil consumption observed in the Canary Islands: 17 000 tonnes in 2002 from 8000 tonnes in 1992, when a new and specific regulation for supplies was implemented (REA, Régimen Específico de Abastecimiento). The REA incentives the import of products not available in the islands, including olive oil²², allowing for an effective nutritional promotion.

Our results support the protective role of olive oil suggested by other studies carried out among Mediterranean populations. Public health recommendations should focus on promoting and facilitating olive oil consumption through political, economical, and social strategies.

Acknowledgements

This study was partially granted by the Spanish Ministry of Health (Fondo de Investigaciones Sanitarias, FIS 98/0917) and the Canary Islands Department of Health (Fundación Canaria de Investigación y Salud, FUNCIS 41/98). Dr García-Segovia's stay at the University of Las Palmas de Gran Canaria was supported by the Research Exchange Program (PPI-00-03) of the Polytechnic University of Valencia.

References

- 1 Willett WC. Dietary fat and breast cancer. In: Willett WC, ed. *Nutritional Epidemiology*, 2nd ed. New York: Oxford University Press, 1998; 377–413.
- 2 Hunter DJ, Spiegelman D, Adami H-O, Beeson L, van den Brandt PA, Folsom AR, *et al.* Cohort studies of fat intake and the risk of breast cancer — a pooled analysis. *New England Journal of Medicine* 1996; **334**: 356–61.
- 3 Holmes MD, Hunter DJ, Colditz GA, Stampfer MJ, Hankinson SE, Speizer FE, *et al.* Association of dietary intake of fat and fatty acids with risk of breast cancer. *Journal of the American Medical Association* 1999; **281**: 914–20.
- 4 Smith-Warner SA, Spiegelman D, Adami H-O, Beeson WL, van den Brandt PA, Folsom AR, *et al.* Types of dietary fat and breast cancer: a pooled analysis of cohort studies. *International Journal of Cancer* 2003; **92**: 767–74.
- 5 Estève J, Kricke A, Ferlay J, Parkin DM. *Facts and Figures of Cancer in the European Community*. Lyon: International Agency for Research on Cancer, 1993.
- 6 Trichopoulou A, Lagiou P, Kuper H, Trichopoulos D. Cancer and Mediterranean dietary traditions. *Cancer Epidemiology, Biomarkers & Prevention* 2000; **9**: 869–73.
- 7 Martín-Moreno JM, Willett WC, Gorgojo L, Banegas JR, Rodríguez-Artalejo F, Fernández-Rodríguez JC, *et al.* Dietary fat, olive oil intake and breast cancer risk. *International Journal of Cancer* 1994; **58**: 774–80.
- 8 Trichopoulou A, Katsouyanni K, Stuver S, Tzala L, Gnardellis C, Rimm E, *et al.* Consumption of olive oil and specific food groups in relation to breast cancer risk in Greece. *Journal of the National Cancer Institute* 1995; **87**: 110–6.
- 9 Morales Suárez-Varela M, Jiménez López MC, Almenar Cubells D, Llopis González A. Effect of the ingestion of food

- and gynecologic risk factors on breast cancer risk in Valencia. *Nutrition Hospitalaria* 1998; **13**: 325–9.
- 10 La Vecchia C, Negri E, Francheschi S, Decarli A, Giacosa A, Lipworth L. Olive oil, other dietary fats, and the risk of breast cancer (Italy). *Cancer Causes & Control* 1995; **6**: 545–50.
 - 11 Trichopoulou A, Lagiou P. *Methodology for the Exploitation of Household Budget Survey Food Data Availability in Six European Countries*. Luxembourg: Office for Official Publications of the European Communities, 1998.
 - 12 Martín-Moreno JM, Boyle P, Gorgojo L, Maisonneuve P, Fernández-Rodríguez JC, Salvini S, *et al.* Development and validation of a food frequency questionnaire in Spain. *International Journal of Epidemiology* 1993; **22**: 512–9.
 - 13 Mataix J, Mañas M, Martínez de Vitoria E. *Tabla de composición de alimentos españoles*, 3rd ed. Granada: Universidad de Granada, 1998.
 - 14 Willett WC. Issues in analysis and presentation of dietary data. In: Willett WC, ed. *Nutritional Epidemiology*, 2nd ed. New York: Oxford University Press, 1998; 321–46.
 - 15 Willett WC, Stampfer MJ. Implications of total energy intake for epidemiologic analysis. In: Willett WC, ed. *Nutritional Epidemiology*, 2nd ed. New York: Oxford University Press, 1998; 273–301.
 - 16 Hunter DJ, Willett WC. Nutrition and breast cancer. *Cancer Causes & Control* 1996; **7**: 56–68.
 - 17 Landa MC, Frago N, Tres A. Diet and the risk of breast cancer in Spain. *European Journal of Cancer Prevention* 1994; **3**: 313–20.
 - 18 Simonsen NR, Fernández-Crehuet Navajas J, Martín-Moreno JM, Strain JJ, Huttunen JK, Martín BC, *et al.* Tissue stores of individual monounsaturated fatty acids and breast cancer: the EURAMIC study. European community multicenter study on antioxidants, myocardial infarction, and breast cancer. *American Journal of Clinical Nutrition* 1998; **68**: 134–41.
 - 19 Caruso D, Berra B, Giavarini F, Cortesi N, Fedeli E, Galli G. Effect of virgin olive oil phenolic compounds on *in vitro* oxidation of human low density lipoproteins. *Nutrition, Metabolism, and Cardiovascular Diseases* 1999; **9**: 102–7.
 - 20 Fitó M, Covas MI, Lamuela-Raventós R, Vila J, Torrents J, Marrugat J. Protective effect of olive oil and its phenolic compounds against low density lipoprotein oxidation. *Lipids* 2000; **35**: 633–8.
 - 21 Prieto-Ramos F, Serra-Majem LL, La Vecchia C, Ramón JM, Tresserras R, Salleras L. Mortality trends and past and current dietary factors of breast cancer in Spain. *European Journal of Epidemiology* 1996; **12**: 141–8.
 - 22 Serra-Majem LL, Cabrera-León A, Sierra-López A. Conclusiones de la Encuesta de Nutrición de Canarias (1997–98). Bases para una política de nutrición en Canarias. *Archivos Latinoamericanos de Nutrición* 2000; **50**(Suppl. 1): 62–70.