#### Environ Health Perspect

### DOI: 10.1289/EHP11391

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### **Supplemental Material**

## Long-Term Exposure to Nitrate and Trihalomethanes in Drinking Water and Prostate Cancer: A Multicase–Control Study in Spain (MCC-Spain)

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# **Table of Contents**

**Table S1.** Characteristics and drinking-water contaminant exposures of the participants excluded from the study population from a Multicase–Control Study in Spain (MCC-Spain) (N=653).

**Table S2.** Mean (standard deviation) of age, adherence to WCRF/AICR cancer prevention score, red and processed meat consumption, fiber intake, total fruit and vegetables intake, vitamin C intake and vitamin E intake across waterborne ingested nitrate tertiles and by case-control status. Multicase–Control Study in Spain (MCC-Spain): 629 prostate cancer cases, 824 controls (N=1,453).

**Figure S1.** Geographic distribution of the water zones with data on nitrate and THM levels included in the study during the exposure window. Multicase–Control Study in Spain (MCC-Spain): 697cases, 927 controls (N= 1,624).

Figure S2. Spearman's rank correlation coefficients between drinking-water exposures.

Additional File- Excel Document

	Controls % or mean (SD)	Cases % or mean (SD)	
	254	200	
n Charles to the	354	299	
Characteristics			
Age (years)	66.4 (9.2)	66.2 (7.7)	
Educational level (%)			
Less than primary	20.3	25.4	
Primary school	31.4	37.1	
Secondary school	26.3	20.4	
University	22.0	17.1	
Family history of prostate cancer (first degree) (%)	15.0	20.7	
Smoking status (%)			
Never	29.9	26.1	
Former	55.4	56.2	
Current smoker	14.7	17.7	
WCRF/AICR cancer prevention score	3.4 (0.9)	3.2 (0.9)	
Intake of red and processed meat (g/day)	78.1 (44.0)	80.5 (52.4)	
Intake of total fiber (g/day)	11.5 (3.9)	10.9 (3.4)	
Intake of fruit and vegetables (g/day)	519 (310)	474 (250)	
Intake of vitamin C (mg/day)	156 (101)	144 (77)	
Intake of vitamin E (mg/day)	11.0 (7.0)	10.5 (6.2)	
Recruitment area (%)			
Asturias	13.6	2.7	
Barcelona	49.7	34.4	
Cantabria	16.1	24.7	
Madrid	16.4	25.4	
Valencia	4.2	12.7	
Drinking-water contaminant exposures			
Average concentrations in residential tap water			
Nitrate (mg/L)	7.4 (4.2)	7.3 (4.2)	
Brominated trihalomethane $(\mu g/L)$	39.1 (36.5)	29 (26.8)	
Chloroform ( $\mu$ g/L)	18.5 (8.9)	18.6 (9.8)	
Daily average waterborne ingestion	· · · ·	~ /	
Nitrate (mg/day)	11 (9.6)	12.2 (10.1)	
Brominated trihalomethane (ug/dav)	25.8 (43.7)	18.5 (34.7)	
Chloroform (µg/day)	15.6 (20.3)	14.6 (23.3)	

**Supplemental Table 1.** Characteristics and drinking-water contaminant exposures of the participants excluded from the study population from a Multicase–Control Study in Spain (MCC-Spain) (N=653)

**Supplemental Table 2.** Mean (standard deviation) of age, adherence to WCRF/AICR cancer prevention score, red and processed meat consumption, fiber intake, total fruit and vegetables intake, vitamin C intake and vitamin E intake across waterborne ingested nitrate tertiles and by case-control status. Multicase–Control Study in Spain (MCC-Spain): 629 prostate cancer cases, 824 controls (N=1,453)\*.

Waterborne ingested nitrate (tertiles)	<5.5 mg/day		5.5–13.8 mg/day		>13.8 mg/day	
	Controls	Cases	Controls	Cases	Controls	Cases
	(n=277)	(n=192)	(n=273)	(n=186)	(n=274)	(n=251)
Age (years)	67.2 (8.3)	65.9 (7.3)	66.3 (8.6)	66.7 (7.5)	66.4 (7.9)	65.5 (7.1)
WCRF/AICR score	3.4 (1.0)	3.3 (0.9)	3.3 (1.0)	3.3 (0.9)	3.3 (1.0)	3.4 (1.0)
Intake of red and processed meat (g/day)	70.2 (36.9)	75.7 (37.6)	73.5 (37.2)	74.3 (39.4)	76.3 (39.6)	80.2 (43.7)
Intake of total fiber (g/day)	11.3 (4.0)	11.4 (3.7)	11.1 (3.7)	11.9 (3.9)	11.4 (3.6)	11.0 (3.2)
Intake of fruit and vegetables (g/day)	483(262)	513 (227)	477 (308)	477 (258)	497 (258)	510 (250)
Intake of vitamin C (mg/day)	149 (83)	154 (77)	147 (105)	146 (84)	147 (86)	151 (80)
Intake of vitamin E (mg/day)	10.0 (4.5)	10.1 (4.5)	10.1 (5.5)	10.4 (5.3)	11.1 (6.2)	11.2 (5.6)

\*These analyses are performed excluding 171 subjects with missing data in the dietary variables.

WCRF/AICR cancer prevention score is based on The World Cancer Research Fund/American Institute for Cancer Research cancer prevention recommendations

**Supplemental Figure 1.** Geographic distribution of the water zones with data on nitrate and THM levels included in the study during the exposure window. Multicase–Control Study in Spain (MCC-Spain): 697cases, 927 controls (N= 1,624)



The intensity of the blue colour represents the number of person-years corresponding to each water zone. Corresponding numeric data can be found in **Excel Table S1**.



**Supplemental Figure 2.** Spearman's rank correlation coefficients between drinking-water exposures