Gazpacho consumption is associated with lower blood pressure and reduced hypertension in a high cardiovascular risk cohort. Cross-sectional study of the PREDIMED trial

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Acronyms: BP, blood pressure; CI, confidence interval; DASH, Dietary Approaches to Stop Hypertension; FFQ, food frequency questionnaire; OR, Odds ratio; PREDIMED, prevention with Mediterranean diet study; SD, standard deviations; TMD, traditional Mediterranean diet.

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Hypertension is a leading cause of death and disableness in the world [1]. Hypertension results from the interaction between genetic and environmental factors. The principal environmental factors influencing blood pressure (BP) are diet, physical activity and psychosocial factors [2]. Changing dietary habits is considered the principal lifestyle measure in BP control [3] since publication of the landmark DASH (Dietary Approaches to Stop Hypertension) trial [4], and the ensuing DASH-Sodium trail with added salt restriction [5]. More recently, increasing epidemiological and clinical evidence points to the traditional Mediterranean diet (TMD) as an alternative dietary pattern for BP control [6,7]. Of note, the healthy diets recommended to prevent hypertension in a high cardiovascular risk cohort. Cross-sectional study of the PREDIMED trial, Nutrition, Metabolism & Cardiovascular Diseases (2012), http://dx.doi.org/10.1016/j.numecd.2012.07.008

**KEYWORDS**
Hypertension; Blood pressure; Gazpacho (Mediterranean vegetable soup); PREDIMED study; Polyphenols

**Abstract**  
Background and aim: Hypertension is a major public health problem and a leading cause of death and disability in both developed and developing countries, affecting one-quarter of the world’s adult population. Our aim was to evaluate whether the consumption of gazpacho, a Mediterranean vegetable-based cold soup rich in phytochemicals, is associated with lower blood pressure (BP) and/or reduced prevalence of hypertension in individuals at high cardiovascular risk.

Methods and results: We selected 3995 individuals (58% women, mean age 67 y) at high cardiovascular risk (81% hypertensive) recruited into the PREDIMED study. BP, weight, and dietary and physical activity data were collected. In multivariate linear regression analyses, after adjusting, moderate and high gazpacho consumption categories were associated with reduced mean systolic BP of 1.9 mm Hg [95% confidence interval (CI): –3.4; –0.6] and –2.6 mm Hg (CI: –4.2; –1.0), respectively, and reduced diastolic BP of –1.5 mm Hg (CI: –2.3; –0.6) and 1.9 mm Hg (CI: –2.8; –1.1). By multiple-adjusted logistic regression analysis, gazpacho consumption was associated with a lower prevalence of hypertension, with OR = 0.85 (CI: 0.73; 0.99) for each 250 g/week increase and OR = 0.73 (CI: 0.55; 0.98) for high gazpacho consumption groups compared to the no-consumption group.

Conclusions: Gazpacho consumption was inversely associated with systolic and diastolic BP and prevalence of hypertension in a cross-sectional Mediterranean population at high cardiovascular risk. The association between gazpacho intake and reduction of BP is probably due to synergy among several bioactive compounds present in the vegetable ingredients used to make the recipe.
Gazpacho intake and reduction of blood pressure

associated with a wide range of health benefits and could be a contributory factor to the low cardiovascular mortality rates observed in southern European countries in comparison with other Western countries [17].

Since the anti-hypertensive food components of gazpacho may act synergistically to further BP reduction, we undertook a substudy of a large nutritional intervention study, the PREDIMED (prevention with Mediterranean diet) trial [18], in order to assess the cross-sectional association between consumption of gazpacho and both BP and prevalent hypertension in individuals at high cardiovascular risk.

Methods

Study subjects

The PREDIMED study is a large, parallel-group, multicenter, randomized, controlled clinical trial of 5-year duration aimed at assessing the effects of the TMD on the primary prevention of cardiovascular disease (www.predimed.org). We undertook a cross-sectional substudy of the baseline data within this larger clinical trial.

The detailed recruitment method and study protocol have been described previously [7,18]. For this substudy, we selected 3995 participants recruited in primary health centers affiliated with the Hospital Clinic of Barcelona, University of Valencia, University of Malaga, Department of Family Medicine (Sevilla), and University of Navarra (Pamplona) in Spain. Eligible participants were community-dwelling men aged 55–80 years and women aged 60–80 years, who were free of cardiovascular disease at baseline and fulfilled at least one of the following two criteria: (1) type-2 diabetes mellitus and/or (2) three or more coronary heart disease (CHD) risk factors: hypertension (systolic BP ≥140 mm Hg or diastolic BP ≥90 mm Hg or treatment with antihypertensive medication), dyslipidemia (LDL cholesterol ≥160 mg/dl; HDL cholesterol ≤40 mg/dl in men or ≤50 mg/dl in women; or under lipid-lowering therapy), obesity or overweight, smoking or family history of early-onset CHD [19]. This trial has been registered with the International Standard Randomised Controlled Trial Number (ISRCTN of London, England) 35739639.

Clinical and dietary measurements

All participants completed at baseline a validated semi-quantitative food frequency questionnaire (FFQ) with 137 items, the validated Spanish version, of the Minnesota Leisure Time Physical Activity Questionnaire, a validated 14-point Mediterranean diet score, and a 47-item questionnaire about education, lifestyle, history of illnesses and medication use. These questionnaires took into account all data during one year before entry into the study, date when BP was measured and hypertension status checked [18]. Trained nurses measured height and weight with a wall-mounted stadiometer and calibrated scales, respectively. BP was measured in triplicate with the participant in a seated position after resting quietly for 5 min, using a validated semi-automatic oscillometer (Omron HEM-705CP [20]; Hoofddorp, The Netherlands) with a 5-min interval between each measurement. The mean of the three systolic and diastolic BP measurements was calculated. Energy and nutrient intake were derived from Spanish food composition tables.

Statistical analyses

Analyses were performed using SPSS software v17.0 (Chicago, USA). Baseline characteristics of the participants were expressed as means or percentages and standard deviations (SD). Variables were examined for normality and skewness (Kolmogorov and Levene tests). ANOVA-one factor was used for analysis of continuous variables and χ²-test for categorical variables. Multivariate linear regression models were used to assess the relationship between systolic and diastolic BP as dependent variables and gazpacho consumption categories as exposure variables (first category: no consumption; second category: moderate consumption, between 1 and 19 g/day; and third category: high consumption, more than 20 g/day), as well as a consumption increase of 250 g/week as an exposure variable, adjusted for potential confounders. Three models were fitted to assess the role of each independent variable (see Table 3). In addition, we carried out a sensitivity analysis of the association of gazpacho consumption with systolic and diastolic BP in a sample restricted to the patients included within the hypertension criterion from the beginning.

Logistic regression models were fitted to assess the relationship between gazpacho consumption as a continuous variable expressed as 250 g/week or gazpacho consumption categories (no consumption, moderate consumption and high consumption) and hypertension status. OR (Odds ratio) and 95% CI (confidence interval) were calculated. P-values <0.05 (two-tailed) were considered significant. Diagnostics for the detection of outliers, multicollinearity, homoscedasticity, and the normality and independence of errors were carried out when appropriate.

Results

We excluded 33 of 3995 eligible participants of the PREDIMED trial for different reasons: not meeting inclusion criteria (n = 15), food allergies (n = 5) or refusing to participate (n = 13). Thus, 3962 participants entered the study, 1626 men [mean age 66.0 (6.5) y] and 2336 women [mean age 67.8 (5.6) y]. Baseline characteristics of the total group by gazpacho consumption category are shown in Table 1. By study design, participants were older than 55 years, mostly overweight [mean body mass index 29.9 (3.6) kg/m²], and with a sizeable burden of cardiovascular risk factors (46.4% diabetics, 81.8% hypertensive, 71.0% with dyslipidemia, 14.8% active smokers and 16.0% with family history of early-onset CHD).

Most participants adhered to a Mediterranean-style diet, with a relatively high consumption of vegetables, fruit, cereals, olive oil and nuts; a moderate intake of legumes, fish, wine, milk and dairy products; and a lower but still relatively high intake of meat, meat products, pastries and dairy products. Table 2 shows the average food consumption of study participants by gazpacho consumption category. The intake of olive oil, total nuts, fruit, vegetables, total fruit and vegetables, legumes, fish or seafood, milk
and dairy products, coffee, chocolate, natural orange juice, fiber, cholesterol, sodium, potassium, total polyphenols, and total energy intake increased across gazpacho consumption categories. Conversely, significant decreasing trends existed for the intake of meat or meat products, cereals and wine. Thus, subjects in the highest gazpacho consumption category reported a significantly greater 14-point Mediterranean diet score than their counterparts.

In multivariate linear regression analyses (Table 3), a significant inverse association between gazpacho consumption category (g/day) and BP was observed. The two upper categories of gazpacho consumption showed lower levels of systolic and diastolic BP than the category no consumption. Adjusted relative differences in systolic BP were $-1.9$ mm Hg for moderate consumption and $-2.6$ mm Hg for high consumption. The relative differences for diastolic BP were $-1.5$ mm Hg for moderate consumption and $-1.9$ mm Hg for high consumption. In this multivariate linear regression analyses, with a consumption increase of 250 g/week as an exposure variable, the adjusted relative differences were $-0.9$ mm Hg and $-0.8$ mm Hg for systolic and diastolic BP, respectively.

In multivariate linear regression analyses of a sample restricted to $\geq 80\%$ of patients included within the hypertension criterion from the beginning, a significant inverse association between gazpacho consumption category (g/day) and BP was observed. No selection bias as described above occurred in this case. Adjusted relative differences in systolic BP were $-2.3$ mm Hg (95% CI: $-4.0$, $-0.7$; $P = 0.006$) for moderate consumption and $-2.5$ mm Hg (95% CI: $-4.3$, $P = 0.006$) for high consumption.

| Table 1 Baseline characteristics of study subjects by categories of gazpacho consumption. |
|----------------------------------|----------------------------------|----------------------------------|----------------|
| Categorization of gazpacho consumption (g/day)$^b$ | No consumption$^b$ (0 g/d) | Moderate consumption (1–19 g/d) | High consumption (>20 g/d) | $P^c$ |
| No. of subjects | 1878 | 994 | 1090 | 0.001 |
| Age (y), mean (SD)$^d$ | 67.56 (6.15) | 66.75 (5.97) | 66.91 (6.29) | 0.004 |
| Women, n (%) | 1099 (58) | 609 (61) | 628 (58) | 0.21 |
| Weight (Kg), mean (SD) | 75.99 (11.45) | 76.54 (11.79) | 77.49 (12.26) | 0.032 |
| BMI$^e$ (kg/m$^2$), mean (SD) | 29.83 (3.69) | 30.09 (3.93) | 30.19 (3.85) | <0.001 |
| Systolic BP$^f$ (mm Hg), mean (SD) | 150 (20) | 148 (20) | 147 (19) | <0.001 |
| Diastolic BP (mmHg), mean (SD) | 84 (11) | 82 (11) | 82 (11) | <0.001 |
| Hypertension, n (%) | 1546 (82) | 819 (82) | 874 (80) | 0.29 |
| Diabetes, n (%) | 843 (45) | 453 (46) | 542 (50) | 0.033 |
| Dyslipidemia, n (%) | 1345 (72) | 713 (72) | 756 (70) | 0.36 |
| Current smoker n (%) | 261 (14) | 146 (15) | 179 (16) | 0.17 |
| Family history of CHD$^g$ n (%) | 261 (14) | 166 (17) | 205 (19) | <0.001 |
| Medication, n (%) | | | | |
| Antihypertensive | 1344 (72) | 714 (73) | 802 (75) | 0.40 |
| Statins or other hypolipidemic drugs | 872 (47) | 479 (48) | 514 (47) | 0.66 |
| Insulin | 109 (6) | 76 (8) | 93 (9) | 0.013 |
| Oral hypoglycemic drugs | 551 (29) | 306 (31) | 398 (37) | <0.001 |
| Aspirin or other antiplatelet drugs | 352 (19) | 222 (22) | 288 (26) | <0.001 |
| Vitamins or supplements, n (%) | 206 (11) | 126 (13) | 133 (13) | 0.32 |
| Educational level, n (%) | | | | |
| Primary school | 1447 (77) | 695 (70) | 766 (70) | <0.001 |
| High school | 248 (13) | 165 (17) | 193 (18) | 0.002 |
| University | 127 (7) | 107 (11) | 109 (10) | <0.001 |
| Energy expenditure in physical activity (kcal/d), mean (SD) | 231.40 (218.15) | 197.39 (200.34) | 202.02 (239.22) | <0.001 |

$^a$ Mean (standard deviation).
$^b$ Reference category.
$^c$ ANOVA-one factor was used for continuous variables and $\chi^2$-test for categorical variables.
$^d$ Standard deviation.
$^e$ Body mass index (calculated as weight in kilograms divided by height in square meters).
$^f$ Blood pressure.
$^g$ Coronary heart disease.
The relative differences for diastolic BP were 1.6 mm Hg (95% CI: 2.5, 0.7; P = 0.001) for moderate consumption and 1.9 mm Hg (95% CI: 2.9, 0.9; P < 0.001) for high consumption.

Logistic regression analysis revealed an inverse association between gazpacho consumption and the prevalence of hypertension (Table 4). When the analysis was adjusted for all possible confounders (model 3), the prevalence of hypertension among gazpacho consumption (250 g/week) was significantly reduced (OR 0.85, 95% CI: 0.73, 0.99; P = 0.04) compared to non-consumption; in addition, the prevalence of hypertension was significantly reduced (OR = 0.73, 95% CI: 0.55; 0.98) for high gazpacho consumption groups compared to the no-consumption group, but it was not significantly reduced when moderate gazpacho consumption group was compared with the no-consumption group.

**Discussion**

In this cross-sectional study of a large Spanish cohort of subjects at high cardiovascular risk, we observed an inverse association between gazpacho consumption, a ready-made Spanish cold vegetable soup rich in carotenoid, vitamin C and polyphenols, and the prevalence of hypertension. In addition, systolic and diastolic BP levels were inversely associated with gazpacho consumption categories and inversely associated with a consumption increase of 250 g/week, after adjustment for potential confounders, including total fruit and vegetable intake. As gazpacho is a mix of vegetables, the results confirm the benefit on BP of increased vegetable intake [3]. The inverse association between gazpacho consumption and BP was independent of total vegetable intake, suggesting synergy in reducing BP.

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Table 3  Multivariate linear regression analyses (difference of means, 95% CI\textsuperscript{a}) of systolic and diastolic blood pressure according to categories of gazpacho consumption and changes associated with a consumption increase of 250 g/week (difference of means, 95% CI).

<table>
<thead>
<tr>
<th>Categories of gazpacho consumption (g/day)</th>
<th>250 g/week</th>
<th>Changes associated with 250 g/week</th>
</tr>
</thead>
<tbody>
<tr>
<td>No consumption\textsuperscript{b} (0 g/d)</td>
<td>150.3</td>
<td>149.4, 151.2</td>
</tr>
<tr>
<td>Moderate consumption (1–19 g/d)</td>
<td>147.7</td>
<td>146.5, 148.9</td>
</tr>
<tr>
<td>High consumption (&gt;20 g/d)</td>
<td>147.5</td>
<td>146.6, 148.6</td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>Systolic blood pressure</th>
<th>250 g/week</th>
<th>Changes associated with 250 g/week</th>
</tr>
</thead>
<tbody>
<tr>
<td>Unadjusted mean</td>
<td>149.4</td>
<td>148.6, 150.1</td>
</tr>
<tr>
<td>Relative differences</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Age-, sex-, weight-adjusted</td>
<td>-2.1</td>
<td>-3.6, -0.6</td>
</tr>
<tr>
<td>Multivariable-adjusted\textsuperscript{d}</td>
<td>-1.9</td>
<td>-3.4, -0.4</td>
</tr>
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</table>

<table>
<thead>
<tr>
<th>Diastolic blood pressure</th>
<th>250 g/week</th>
<th>Changes associated with 250 g/week</th>
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</thead>
<tbody>
<tr>
<td>Unadjusted mean</td>
<td>83.6</td>
<td>83.2, 84.1</td>
</tr>
<tr>
<td>Relative differences</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Age-, sex-, weight-adjusted</td>
<td>-1.5</td>
<td>-2.3, -0.7</td>
</tr>
<tr>
<td>Multivariable-adjusted\textsuperscript{d}</td>
<td>-1.5</td>
<td>-2.3, -0.6</td>
</tr>
</tbody>
</table>

\textsuperscript{a} Confidence intervals.
\textsuperscript{b} Reference category.
\textsuperscript{c} \textit{P}: two-sided test of significance.
\textsuperscript{d} Adjusted for sex, age, weight, smoking status, energy expenditure in physical activity, educational level, medication intake (anti-hypertensive drugs, statins or other hypolipidemic drugs, insulin, oral hypoglycemic drugs, aspirin or other antiplatelet drug), supplements taken in the last month, dyslipidemia, family history of coronary heart disease, diabetes, as well as for intake of carbohydrates, total fruit and vegetables, alcohol, sodium, potassium, 14-unit Mediterranean diet score and total energy intake.
among bioactive compounds in a recipe that includes a complex mixture of plant foods, with olive oil that increase bioavailability of phenolic compounds [21].

Hypertension is an unequivocal risk factor for cardiovascular morbidity and mortality, and is the main risk factor for stroke in both men and women. The estimated total number of adults with hypertension in 2000 was 972 million (957–987 million) and is predicted to increase by about 60% to a total of 1.56 billion in 2025 [22].

The first step in the prevention and management of hypertension is to follow a plant-based diet such as the DASH or Mediterranean diets [3], which are considered rich sources of phytochemicals and are inversely associated with high BP in epidemiologic studies [4].

Recently, "gazpacho" has acquired importance in Western societies due to the increasing interest in the Mediterranean diet. A serving of "gazpacho" (250 g) typically contributes 120 kcal, which comes mainly from olive oil. The mixture of all the ingredients contributes to the intake of carbohydrates (40 g/kg) and fibers (12 g/kg). Besides, gazpacho contains protein (10 g/kg), and sizeable amounts of vitamins and minerals. The sodium content in "gazpacho" is high compared to that of plant foods due to addition of salt in the recipe, the mean value founded in "gazpacho" is 1,073 mg/kg [23].

Recent research has shown that the raw ingredients of gazpacho exert potentially valuable antihypertensive effects. Thus, a diet rich in fruit and vegetables may help prevent BP increases and reduce elevated BP levels [24]. In the Mediterranean cohort of the Seguimiento Universidad de Navarra (SUN) study [10], the prevalence of hypertension was inversely associated with fruit and vegetable consumption. In a study of 983 Indian adults, increasing fruit and vegetable intake was inversely associated with systolic BP [25].

Tomato is a critical ingredient of gazpacho, and tomato consumption has been associated with reduced cardiovascular risk factors. Tomato extract supplementation was associated with a clinically significant reduction of systolic and diastolic BP in hypertensive patients [15]. Garlic extract has also been associated with reduced systolic BP in patients with hypertension [16]. Mediterranean diets enriched in olive oil have been associated with reduced systolic and diastolic BP in subjects with overweight [6], or increased cardiovascular risk [7]. Finally, a significant decrease in systolic BP was observed in 24 hypertensive patients after 1-wk treatment with a daily dose of 4 capsules of an onion–olive oil maceration product [26]. Thus, the main ingredients of gazpacho have been associated with reduced BP in epidemiological and clinical studies.

High salt intake harms the cardiovascular system by raising BP and might be directly responsible for increasing left ventricular mass, as suggested in clinical and experimental studies [27,28]. Consistent with this, there is also evidence that high salt intake aggravates and, inversely, dietary salt restriction prevents (or at least mitigates) left ventricular hypertrophy in patients with essential hypertension. Some clinical trials have shown that reducing the sodium chloride content of typical diets in the United States or northern Europe lowers BP, and guidelines recommend reducing the daily dietary sodium intake to 100 mmol (equivalent to 2.3 g of sodium or 5.8 g of sodium chloride) or less.

The low-sodium DASH diet produced greater reductions in systolic and diastolic BP than either the DASH diet alone or a reduction in sodium alone, compared with the high-sodium control diet [5]. In a meta-regression analysis of randomized trials [8], a median –77 mmol/24 h sodium reduction was associated with a change of –2.54 mm Hg and –1.96 mm Hg in systolic and diastolic BP, respectively.

In spite of the salt content of homemade and commercial gazpachos, its consumption in the current study was associated with a significant reduction in systolic and diastolic BP. Presumably, the sodium content of gazpacho is counteracted by the high levels of potentially antihypertensive minerals, such as potassium, magnesium, and calcium [3], contained in its main components. Part of the anti-hypertensive effect of gazpacho may also be attributed to its phenolic content. The main sources of polyphenols are fruit, vegetables and beverages such as wine, tea and also gazpacho. By example, the total phenol content of 12 tested commercial Mediterranean gazpachos ranged between 11.54 and 17.52 mg/100 g [29].

Urinary polyphenol excretion, a biomarker of total polyphenol intake, was directly related to fruit and vegetable intake and inversely associated with BP levels and the prevalence of hypertension in another cross-sectional study of the PREDIMED trial [24]. Thus, clinical and epidemiologic evidence supports the antihypertensive effect of

Table 4 Multivariate adjusted odds ratios (95% CI) for hypertension by gazpacho consumption expressed as 250 g/week and as categories of gazpacho consumption (moderate and high consumption groups compared to the no-consumption group).

<table>
<thead>
<tr>
<th>Model</th>
<th>ORb</th>
<th>95% CI for OR</th>
<th>Pd</th>
</tr>
</thead>
<tbody>
<tr>
<td>1 For 250 g/week</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Model 1</td>
<td>0.93</td>
<td>0.84</td>
<td>1.02</td>
</tr>
<tr>
<td>Model 2</td>
<td>0.92</td>
<td>0.84</td>
<td>1.01</td>
</tr>
<tr>
<td>Model 3</td>
<td>0.85</td>
<td>0.73</td>
<td>0.99</td>
</tr>
<tr>
<td>Moderate</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>compared</td>
<td>0.71</td>
<td>0.63</td>
<td>0.81</td>
</tr>
<tr>
<td>with no-consumption</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>High</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>compared</td>
<td>0.80</td>
<td>0.73</td>
<td>0.87</td>
</tr>
<tr>
<td>with no-consumption</td>
<td></td>
<td></td>
<td></td>
</tr>
</tbody>
</table>

CI: Confidence intervals. OR: Odds ratio. a: Model 1, unadjusted; Model 2 was adjusted for sex, age and weight; Model 3 adjusted as in Model 2 plus for smoking status, physical activity, educational level, energy expenditure in physical activity and, medication intake (antihypertensive drugs, statins or other hypolipidemic drugs, insulin, oral hypoglycemic drugs, aspirin or other antiplatelet drug), supplements taken in the last month, dyslipidemia, family history of coronary heart disease, diabetes, as well as for intake of carbohydrates, total fruit and vegetables, alcohol, sodium, potassium, 14-unit Mediterranean diet score and total energy intake. b: Two-sided test of significance.
foods and diets rich in polyphenols. Because greater gazpacho intake means higher polyphenol consumption, we suggest that the observed inverse association between gazpacho consumption categories and BP relates in part to a beneficial effect of polyphenols intake on BP.

Our study has limitations. First, given that our study subjects were elderly and at high risk of cardiovascular disease, the results may not be generalized to other populations. However, since most study subjects were hypertensive, our findings confirm the usefulness of polyphenol-rich diets in the management of hypertension. Another limitation of our study is its cross-sectional nature, which does not allow establishing definite causal relationships. Nevertheless, reverse causality bias seems improbable, since gazpacho contains salt and its regular use is not recommended to patients with hypertension. Clearly, clinical trials aimed at exploring the effects of gazpacho consumption on BP are warranted.

In conclusion, in this cross-sectional study within the framework of the PREDIMED trial in a high-risk Mediterranean population, gazpacho consumption was inversely associated with BP levels and prevalence of hypertension. However, the observation that systolic and diastolic BP decrease with increasing gazpacho consumption should be confirmed in randomized clinical trials. Further studies are also needed to clarify whether functional compounds present in gazpacho, differently contribute to this apparent beneficial health effect, counteracting the possible deleterious effect of the salt that it contains.

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Appendix 1. Other PREDIMED Investigators

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