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## Body Mass Index and Nutrition as Determinants of Health and Disease in Population of Croatian Adriatic Islands

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**Aim** To investigate the relationships between body mass index (BMI), dietary habits, and cardiovascular risk factors in isolated Adriatic island populations of Croatia.

**Methods** Random sample of subjects (n = 1001) was interviewed, using a validated questionnaire developed for this research program. Dietary habits were assessed on the basis of applied Food Frequency Questionnaire (FFQ). Biochemical analyses of total cholesterol, high density lipoprotein (HDL), low density lipoprotein (LDL), triglycerides, and blood glucose were performed. Blood pressure (mm Hg), height (m), and weight (kg) were measured following standard procedures.

**Results** Out of 1001 study participants, 507 (50.7%) were overweight (247 [48.7%] men and 260 [51.3%] women), whereas 268 (26.8%) were obese (122 [45.5%] men and 146 [54.5%] women). In both genders, there was a positive correlation between body mass index (BMI) and levels of triglycerides ( $P < 0.001$ ), glucose ( $P < 0.001$ ), diastolic blood pressure ( $P < 0.001$ ), and systolic blood pressure (male:  $P = 0.002$ , female:  $P < 0.001$ ). Logistic regression showed that female gender (OR, 2.31; 95% CI, 1.61-3.31), overweight (OR, 1.97; 95% CI, 1.34-2.88), obesity (OR, 1.90; 95% CI, 1.22-2.96), more frequent consumption of meat (OR 1.17; 95% CI, 1.06-1.30) and beer (OR 1.14; 95% CI, 1.03-1.27), and less frequent consumption of potatoes (OR 0.91; 95% CI, 0.83-0.99) were predictive for the presence of cardiovascular risk factors.

**Conclusion** Prevalence of obesity and related health outcomes was surprisingly high for the studied population. We found a correlation between BMI, dietary habits, and cardiovascular risk factors.

The health of an individual and the population in general is the result of interactions between genetics and a number of environmental factors. An environmental factor of major importance is nutrition (1), which is also one of the key determinants of health care expenditures (2).

Excessive body weight is a major problem in industrialized and developed countries where it has reached the proportions of an epidemic (3). Overweight and obesity have been related to increased morbidity and mortality rates due to diabetes mellitus, several forms of cancer, digestive disease, and coronary heart diseases (4,5).

Furthermore, overweight and obesity are believed to represent an independent risk for cardiovascular diseases. A part of the increased risk of cardiovascular diseases conferred by increased body weight is explained by the effects on blood pressure, glucose tolerance, and plasma lipid metabolism (6,7). Because of the strong relationship between excessive body weight and these diseases, it is important to provide information on the determinants of overweight and obesity from population-based surveys (6,7).

Body mass index (BMI) is an easily and reliably obtained measure of relative body size. It is often used as an indirect index of adiposity and has been strongly associated with cardiovascular disease risk (8).

During the few last years, cardiovascular diseases have caused approximately 4.3 million deaths in Europe and 1.9 million deaths in the European Union (EU) per year, representing 49% of all deaths in Europe and 42% in the EU (9). In Croatia, these diseases also represent a leading cause of mortality and morbidity: the standardized death rate has been increasing for the last three decades, and reached 91.7 per 100 000 population in 2001 (10).

At the same time, in Finland, Sweden, Norway, and Denmark, the mortality rates of standardized deaths rates from coronary heart disease have been declining at all ages over the past decades (11,12). The reasons for the de-

cline are probably healthier lifestyle, reduction of risk factors, and better treatment of diseases. Populations in South European countries, such as France, Spain, and Portugal, have relatively low death rates due to the Mediterranean diet (13).

Studies have suggested that health-conferring benefits of the Mediterranean diet are mainly due to high consumption of fish, fruit, and vegetables. More recent research has focused on other important factors, such as olives and olive oil. Fiber, especially from wholegrain-derived products, fruit, and vegetables, is an important source of dietary antioxidants (14). Furthermore, data suggest that the traditional Mediterranean dietary pattern is inversely associated with body mass index (BMI) and obesity (15).

The aim of the study was to investigate the relationship between BMI and dietary habits and cardiovascular risk factors in isolated Adriatic island populations of Croatia.

## Subjects and Methods

### Subjects

This study is the part of the "1001 Dalmatians" research program, which was launched with an aim to study isolated populations of Croatian islands (Rab, Vis, Lastovo, and Mljet) that share similar environment and lifestyle (16). A random sample of 1001 adult inhabitants was selected from the total number of inhabitants of these 4 islands. Sampling was based on computerized randomization of the most complete and accessible population registries in each village, which included medical records (Mljet and Lastovo islands), voting lists (Vis island), and household numbers (Rab island). The field work of data gathering was performed during 2002 and 2003 by the team of researchers from the Andrija Štampar School of Public Health, Zagreb University School of Medicine, and the Institute for Anthropological Research in Zagreb, Croatia.

The ethical approval for this research was obtained from the Research Ethics Committee of the Zagreb University School of Medicine, Zagreb, Croatia. Informed written consent was obtained from all participants in the study.

#### **Measures of health-related variables**

All of the examinees were first interviewed by one of the trained surveyors, using a validated questionnaire developed specifically for this research program. The details of this program are described elsewhere (17). The questionnaire was designed for collecting extensive information on personal data (name, date and place of birth, gender, marital status, occupation, and various lifestyle variables), health complaints, drug intake, and hospitalization records. It also included Food Frequency Questionnaire (18). The consumption frequency of the selected foodstuffs (dietary habits) was determined on the basis of the reported weekly frequency of intake: 5-7 times a week, 3-4 times a week, 1-2 times a week, or never.

Height and body mass were measured using a single anthropometer (Hospitalija, Zagreb, Croatia). Body mass index was determined as weight divided by height squared ( $\text{kg}/\text{m}^2$ ). Subjects were classified into three groups, as follows: normal (acceptable) weight,  $\text{BMI} < 25$ ; overweight,  $\text{BMI} 25-29.9$ ; obesity,  $\text{BMI} \geq 30$  (3). Since no underweighted persons were identified, there was no need to form this group.

Blood pressure (mm Hg), height (m), and weight (kg) were each measured by a single observer in local health centers and dispensaries between 8 and 11 AM, following the standard procedure (19). Blood pressure was measured on the right forearm in sitting position. Two measurements of both systolic and diastolic blood pressure in each individual were taken two times in a 5-minute interval. The mean value obtained from two readings was used in the analysis. Hypertension was defined according to World Health Organization (WHO) criteria as systolic

blood pressure  $\geq 140$  mm Hg or diastolic blood pressure  $\geq 90$  mm Hg (20).

Biochemical analyses of high and low density lipoproteins (HDL and LDL) and total cholesterol, triglycerides, and blood glucose were performed from fasting blood samples taken from the examinees between 7 and 9 AM. Plasma and serum were rapidly frozen and stored at  $-20^\circ\text{C}$  in 200 ml aliquots using standardized sample handling procedures. They were then transported frozen within a maximum of 3 days to the single biochemical laboratory, based in Zagreb. The laboratory was chosen as it has been internationally accredited for performing this type of analysis and included in the internal quality assessment by Roche and Olympus, and in external monitoring programs by Croatian reference center for biochemical measurements and RIQAS international agency for quality control (21).

#### **Statistical analysis**

Descriptive statistics were given as median with range, and with absolute and relative frequencies. Kruskal-Wallis test was used for multiple group comparisons (lipids, glucose, and blood pressure values according to BMI groups), with *post hoc* Mann-Whitney test for significant differences. Correlations of BMI, laboratory findings, and blood pressures were evaluated using Spearman rank correlation coefficients due to non-normal distributions of most values. Differences between qualitatively expressed originally quantitative variables regarding BMI groups were tested with  $\chi^2$  test. Logistic regression was used to estimate prediction (power) of dietary habits, age, BMI, and gender for presence of cardiovascular risk factors (defined as presence of high cholesterol, and/or high LDL, and/or low HDL levels). Statistical significance of Spearman rank correlation coefficients of biochemical variables, blood pressure, and BMI was used as a criterion for defining the presence of cardiovascular risk. The maximal acceptable significant level was  $\alpha = 0.05$ . All statistical procedures were performed using

STATISTICA, version 7.1 (StatSoft, Inc., Tulsa, OK, USA).

## Results

We examined 454 men (45.4%), with the mean age of  $55.0 \pm 15.4$  years (range 18-87) and 547 women (54.6%), with the mean age of  $55.0 \pm 14.0$  (range 19-88) years. Among them, 247 (54.4%) men and 260 (47.5%) women were overweight and 122 (26.9%) men and 146 (26.7%) women were obese.

In both genders, significant differences were found among normal, overweight, and obese examinees in the levels of triglycerides, glucose, diastolic blood pressure (for both genders  $P < 0.001$ ), total serum cholesterol (men,  $P = 0.002$ ; women,  $P = 0.013$ ), and systolic blood pressure (men,  $P = 0.030$ ; women,  $P < 0.001$ ; Table 1). For LDL levels, the difference between BMI groups was significant in men only ( $P < 0.001$ ).

Overweight men had significantly higher levels of cholesterol, LDL, triglycerides, glucose, and diastolic pressure than men of normal weight. Obese men had significantly higher levels of cholesterol, triglycerides, glucose, systolic and diastolic pressure than men of normal weight. Obese men had significantly higher levels of triglycerides, glucose, systolic and diastolic pressure, when compared with overweight men.

Overweight women had significantly higher levels of cholesterol, triglycerides, systolic and diastolic pressure than women of normal weight. Obese women had significantly higher levels of cholesterol, triglycerides, glucose, systolic and diastolic pressure compared with women of normal weight. When overweight and obese women were compared, obese women had significantly higher levels of glucose, systolic and diastolic pressure. All other relevant variables showed no significant differences between groups (Table 1).

The differences between BMI groups were also found when values of cardiovascular risk factors were dichotomized according to referent values (Table 2). Abnormal values of all observed risk factors predominated in the overweight group. For all risk factors, the distribution of BMI groups was significant on at least 5% level.

As far as dietary habits are concerned, the majority of participants consumed vegetables and fruit ( $n = 758$ ; 75.8%), olive oil ( $n = 638$ ; 63.8%), potatoes ( $n = 422$ ; 42.2%), and wine ( $n = 409$ ; 40.9%) frequently, ie, 5 to 7 times a week. Fish was consumed 5 to 7 times a week by 98 (9.8%) subjects and 3 to 4 times a week by 345 (34.5%) subjects. Meat and milk was mostly consumed 3 to 4 times a week. Rice and pasta was mostly consumed 1 to 2 times a week. Beer was consumed 5-7 times a week by 6.9% ( $n = 69$ ) participants,

**Table 1.** Cardiovascular risk factors (median, range) by gender and body mass index (BMI) in the population of Croatian Adriatic islands

| Parameters <sup>‡</sup>          | BMI groups*               |                         |                      |                       |                            |                         |                      |                       |
|----------------------------------|---------------------------|-------------------------|----------------------|-----------------------|----------------------------|-------------------------|----------------------|-----------------------|
|                                  | men (n = 454)             |                         |                      |                       | women (n = 547)            |                         |                      |                       |
|                                  | normal weight<br>(n = 85) | overweight<br>(n = 247) | obesity<br>(n = 122) | <i>P</i> <sup>†</sup> | normal weight<br>(n = 141) | overweight<br>(n = 260) | obesity<br>(n = 146) | <i>P</i> <sup>†</sup> |
| Total cholesterol (<5.2 mmol/L)  | 5.2 (3.12-9.78)           | 5.9 (3.30-11.41)        | 5.4 (3.18-9.99)      | 0.002                 | 5.7 (2.92-11.72)           | 6.2 (3.71-11.75)        | 6.2 (3.52-10.44)     | 0.013                 |
| LDL cholesterol (2.0-3.9 mmol/L) | 3.6 (1.42-8.38)           | 4.1 (1.45-8.51)         | 3.5 (1.58-7.34)      | <0.001                | 4.1 (1.42-9.87)            | 4.4 (1.89-9.72)         | 4.4 (1.34-7.62)      | 0.273                 |
| HDL cholesterol (0.9-1.4 mmol/L) | 1.1 (0.59-1.31)           | 1.0 (0.50-1.31)         | 1.1 (0.55-1.41)      | 0.224                 | 1.1 (0.77-1.45)            | 1.0 (0.62-1.92)         | 1.1 (0.51-1.34)      | 0.095                 |
| Triglycerides (0-1.5 mmol/L)     | 1.1 (0.46-7.07)           | 1.2 (0.46-10.61)        | 1.4 (0.61-11.79)     | <0.001                | 1.1 (0.44-3.15)            | 1.2 (0.49-4.25)         | 1.3 (0.58-5.17)      | <0.001                |
| Glucose (4.2-6.4 mmol/L)         | 5.3 (3.73-8.58)           | 5.4 (3.78-18.15)        | 5.8 (4.02-16.24)     | <0.001                | 5.2 (3.54-13.58)           | 5.4 (3.53-18.68)        | 5.6 (3.73-20.86)     | <0.001                |
| Systolic pressure (<140 mm Hg)   | 141.2 (94-200)            | 145.6 (100-198)         | 147.2 (106-233)      | 0.030                 | 135.5 (99-218)             | 138.6 (97-207)          | 149.3 (95-227)       | <0.001                |
| Diastolic pressure (<90 mm Hg)   | 86.5 (63-123)             | 89.7 (55-120)           | 92.8 (61-124)        | <0.001                | 83.9 (50-121)              | 87.3 (55-125)           | 91.1 (68-126)        | <0.001                |

\*Normal weight - BMI < 25; overweight - BMI = 25-29.99; obesity - BMI ≥ 30. There were no underweight persons.

†Kruskal-Wallis test.

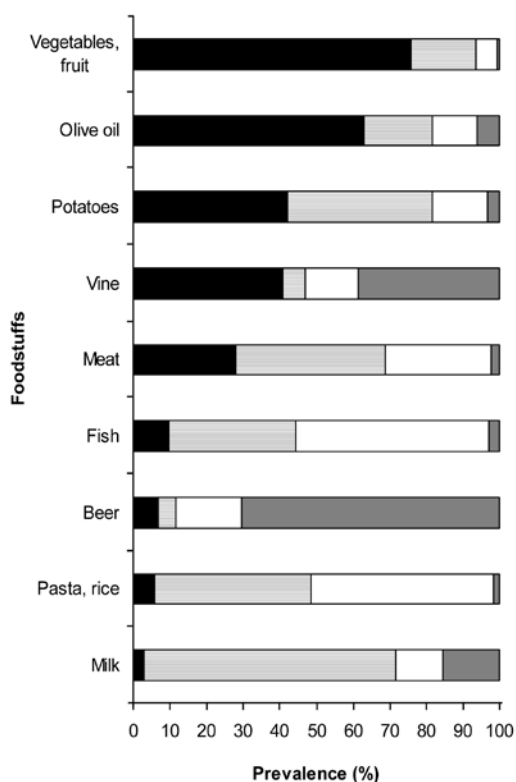
‡The value in the brackets is laboratory referent value.

**Table 2.** Blood pressure and concentration of lipids and glucose, according to referent values by body mass index (BMI)

| Parameters                  | BMI groups*             |                      |                   | total n = 1001 | P <sup>†</sup> |
|-----------------------------|-------------------------|----------------------|-------------------|----------------|----------------|
|                             | normal weight (n = 226) | overweight (n = 507) | obesity (n = 268) |                |                |
| Total cholesterol (mmol/L): |                         |                      |                   |                |                |
| <5.2                        | 79 (28.1)               | 125 (44.5)           | 77 (27.4)         | 281 (100.0)    | 0.016          |
| ≥5.2                        | 147 (20.4)              | 382 (53.0)           | 191 (26.6)        | 720 (100.0)    |                |
| LDL cholesterol (mmol/L):   |                         |                      |                   |                |                |
| ≤3.9                        | 109 (25.3)              | 194 (45.1)           | 127 (29.5)        | 430 (100.0)    | 0.010          |
| >3.9                        | 117 (20.5)              | 313 (54.8)           | 141 (24.7)        | 571 (100.0)    |                |
| HDL cholesterol (mmol/L):   |                         |                      |                   |                |                |
| ≥0.9                        | 203 (24.5)              | 410 (49.4)           | 217 (26.1)        | 830 (100.0)    | 0.007          |
| <0.9                        | 23 (13.5)               | 97 (56.7)            | 51 (29.8)         | 171 (100.0)    |                |
| Triglycerides (mmol/L):     |                         |                      |                   |                |                |
| ≤1.5                        | 184 (26.7)              | 349 (50.7)           | 156 (22.6)        | 689 (100.0)    | <0.001         |
| >1.5                        | 42 (13.5)               | 158 (50.6)           | 112 (35.9)        | 312 (100.0)    |                |
| Glucose (mmol/L):           |                         |                      |                   |                |                |
| 4.2-6.4                     | 209 (24.3)              | 443 (51.5)           | 209 (24.3)        | 861 (100.0)    | <0.001         |
| >6.4                        | 17 (12.1)               | 64 (45.7)            | 59 (42.1)         | 140 (100.0)    |                |
| Systolic pressure (mm Hg):  |                         |                      |                   |                |                |
| <140                        | 122 (27.1)              | 234 (52.0)           | 94 (20.9)         | 450 (100.0)    | <0.001         |
| ≥140                        | 104 (18.9)              | 273 (49.5)           | 174 (31.7)        | 551 (100.0)    |                |
| Diastolic pressure (mm Hg): |                         |                      |                   |                |                |
| <90                         | 157 (28.1)              | 283 (50.7)           | 118 (21.1)        | 558 (100.0)    | <0.001         |
| ≥90                         | 69 (15.6)               | 224 (50.6)           | 150 (33.9)        | 443 (100.0)    |                |

\*Normal weight – BMI<25; overweight – BMI = 25-29.99; obesity – BMI≥30. There were no underweight persons.

† $\chi^2$  test.



**Figure 1.** Dietary habits of the population of Croatian Adriatic islands (n=1001), expressed as the frequency of consumption of select foodstuffs, were determined on the basis of reporting the weekly frequency of intake. Closed bar, 5-7 times a week; dark grey bar, 3-4 times a week; open bar, 1-2 times a week; grey bar, never.

whereas 70.3% (n = 704) of subjects never drank beer (Figure 1).

Due to the fact that triglycerides, glucose, and blood pressure showed highly significant correlation with BMI, they were not included in the definition of “being ill” (presence of cardiovascular risk factors), which was the criterion variable in the logistic regression analysis.

Logistic regression showed that female gender (OR, 2.31; 95% CI, 1.61-3.31), overweight (OR, 1.97; 95% CI, 1.34-2.88), obesity (OR, 1.90; 95% CI, 1.22-2.96), more frequent consumption of meat (OR, 1.17; 95% CI, 1.06-1.30) or beer (OR, 1.14; 95% CI, 1.03-1.27), and less frequent consumption of potatoes (OR=0.91; 95% CI 0.83-0.99) were predictive for presence of cardiovascular risk factors, defined as high cholesterol, and/or high LDL, and/or low HDL levels (Table 3).

## Discussion

Our study demonstrated the correlation between BMI and dietary habits in relation to cardiovascular risk factors. Differences between BMI



**Table 3.** Effects of dietary habits, gender, age, and body mass index (BMI) on presence of risk factors\*

| Variable              | OR (95% CI)      |
|-----------------------|------------------|
| Female gender         | 2.31 (1.61-3.31) |
| Overweight            | 1.97 (1.34-2.88) |
| Obesity               | 1.90 (1.22-2.96) |
| Age                   | 1.01 (0.99-1.02) |
| Fish                  | 0.99 (0.88-1.11) |
| Meat                  | 1.17 (1.06-1.30) |
| Vegetables and fruits | 1.04 (0.99-1.09) |
| Milk                  | 0.97 (0.92-1.02) |
| Pasta and rice        | 0.94 (0.88-1.01) |
| Potatoes              | 0.91 (0.83-0.99) |
| Olive oil             | 1.06 (0.99-1.14) |
| Vine                  | 1.02 (0.97-1.08) |
| Beer                  | 1.14 (1.03-1.27) |

\*Logistic regression. Dependent variable included presence of high cholesterol and/or high LDL and/or low HDL levels.

groups (normal, overweight, and obese) were found for all parameters tested, except HDL values in both genders and LDL values in women. Also, statistically significant differences in the distribution of qualitatively transformed cardiovascular risk factors, regarding referent values between BMI groups were found. Logistic regression showed that female gender, overweight, obesity, and frequent consumption of meat or beer had statistically significant positive predictive value for the classification of "being ill." On the other hand, frequent consumption of potatoes showed statistically significant protective value for "being ill."

Female gender, overweight, and obesity had OR between 2.31 to 1.90. There was 17% higher probability to be classified as "ill" for those who frequently consumed meat, whereas there was only 9% smaller probability for "being ill" for frequent potato eaters. Overweight subjects showed pathological values of total cholesterol, LDL, and HDL (53.0%, 54.8%, and 56.7% respectively).

These results are in accordance with the results of other studies, which showed that overweight and obesity are consistent parameters associated with cardiovascular risk in most populations (6,22-25).

Our findings suggest that female gender, overweight, and obesity are predictors of cardiovascular risk, and are very consistent in their relationships. This could partly be related to the age

of women. The average age of the surveyed population was  $55.0 \pm 14.8$  (range: 18-88 years), with most of the women aged 50 years and older. This is menopausal age, which includes hormonal deficiency associated with higher level of biochemical parameters, and can contribute to higher cardiovascular risk (26). Also, CRO-CAN study conducted on Croatian population showed that the prevalence of cardiovascular risk factors was generally high, but the hierarchy varied between regions and between sexes (27).

We also wanted to find out whether there was a correlation between cardiovascular risk factors and dietary habits of an isolated island population. Most of the authors analyzed the influence of the Mediterranean diet on the decrease of BMI and thus, indirectly, on cardiovascular risk factors (15,28).

Although the islands and coast are home to more elderly population, the mortality from circulatory diseases is lower. This means that among other factors, the Mediterranean diet plays an important role in preventing atherosclerosis, and decreasing the morbidity and mortality from circulatory diseases in Croatia (29).

As proposed by other authors (15), we expected to confirm the beneficial impact of the Mediterranean diet on cardiovascular status of the surveyed island population. Although the investigated island population showed significant frequency of consuming foodstuffs characteristic for the Mediterranean diet (vegetables and fruit, olive oil, fish, and vine), none of these foodstuffs was found to be a negative predictor for cardiovascular risk. This could suggest that the intake of typical Mediterranean foodstuffs is decreasing, whereas the intake of meat is on the increase. The change in dietary pattern is partly related to the change in lifestyle which occurred due to socioeconomic factors, existing food markets, and the wide range of industrially prepared foodstuffs available. This change has been facilitated by tourism which has substituted agriculture as the main source of income, as well as by better con-

nections with the mainland which enabled better supply with meat and industrially prepared food.

Similar changes were noticed by Ortega et al in the dietary pattern of a Spanish population (30). They reported that in the last few years, changes in lifestyle have triggered a move away from the recommended intakes of several nutrients. The intake of cereals and vegetables has decreased, whereas the intake of fats and animal products has increased.

Our findings are similar to the results of the First Croatian Health survey, conducted in 1995-1997. The survey showed that olive oil consumption has decreased due to high market price and low local production, whereas the consumption of red meat has increased. Also, there were no regional differences between continental and coastal region in the consumption of meat and eggs. Obviously, some changes in the structure of diet and dietary habits have occurred over time, making the dietary habits on the islands more similar to the continental diet pattern. The change may have multi factorial etiology such as personal attitudes, change in tradition and economic factors, as well as availability of industrial food products, as opposed to private individual food production (31,32).

Our study showed high prevalence of obesity and cardiovascular risk factors in isolated island populations of the Mediterranean. According to some authors, substantial reassessment of the importance of obesity for cardiovascular disease in most regions of the world is needed (33). Also, in order to obtain better insight into the nutrition quality we recommend a 24-hour recall study in order to provide quantitative data and meal pattern.

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## ANNOUNCEMENT

**XII. MEETING OF THE EUROPEAN ASSOCIATION FOR HAEMATOPATHOLOGY**

October 8-11, 2006, Vienna/Austria

**Main Topic:**

Small B-cell lymphomas

**Congress Secretariat:**

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