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Source / Izvornik: **Collegium Antropologicum, 2012, 36, 183 - 188**

Journal article, Published version

Rad u časopisu, Objavljena verzija rada (izdavačev PDF)

Permanent link / Trajna poveznica: <https://um.nsk.hr/um:nbn:hr:105:173789>

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Download date / Datum preuzimanja: **2024-12-21**



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Association of Elevated Body Mass Index and Hypertension with Mortality: the CroHort Study

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ABSTRACT

Aim of this study was to investigate association of elevated body mass index and hypertension with general mortality in the cohort from Croatian Adult Health Cohort Study (CroHort). Risk of death according to body mass index (BMI) and blood pressure category, in period 2003/08, was calculated for 7,490 respondents, out of which 6,682 were alive and 808 were dead in 2008. Among men aged 65 and more, elevated BMI was associated with lower risk of death, in accordance with obesity paradox recorded in some previous studies which showed that older men with higher BMI have lower risk of death. Among women aged 50–64 years, being hypertensive was associated with increased risk of death compared to normotensive respondents. Despite obesity paradox which should be further explored, activities on primary and secondary prevention of excess weight and hypertension should be encouraged as means to prevent premature mortality in Croatian population.

Key words: body mass index, overweight, obesity, hypertension, mortality, CroHort study

Introduction

Cardiovascular diseases are the leading cause of death in developed countries and are a rapidly growing cause of death in low- and middle-income countries. According to the Global Burden of Disease Study in 2002, the leading cause of burden of disease in Europe are cardiovascular diseases with a 23% share in all DALYs. Croatia has, according to the same source, cardiovascular diseases at the top of the list, with a proportion of 27% of all DALYs.

Over half of burden of disease are caused by seven leading risk factors. The leading risk factor globally is high blood pressure with a share of 12.8% and overweight is ranked 5th with a share of 7.8% in the DALY burden. Among Croatians, elevated blood pressure is ranked second (after smoking) with a share of 13.8%¹.

In Croatia, ca. 50,000 people die annually. Mortality rate is relatively stable at around 11‰. More than 50% of all death causes are related to circulation diseases, due to which 26,235 people died (591.2/100,000). The leading diagnostic entities are ischemic heart disease and cerebrovascular diseases (shares of 39% and 31% in circulatory disease group)². Ischemic heart disease caused death of 10,542 people (standardized rates: 157.75 / 100,000)

which ranked Croatia in European countries with medium high mortality rates³.

High blood pressure and obesity are recognized as important risk factors for mortality all around the world^{4,5}. Each year more than 7 million people die as a result of raised blood pressure and 2.6 million people as a result of being overweight or obese^{6–8}.

The studies based on the Croatian Health Survey 2003 data noted prevalence of arterial hypertension (therapy administration or systolic pressure=140, or diastolic pressure=90) of 45.6% for man and 43% for women⁹.

In Croatia, the prevalence of overweight and obesity has been estimated at almost 60% of the adult population¹⁰, and any association between overweight and mortality might have important clinical and public health implications. Overweight and obese people have an increased risk of developing many serious conditions e.g. cardiovascular disease^{11,12}, diabetes mellitus¹³, certain cancers¹⁴ like endometrial¹⁵, ovarian¹⁶, breast^{16,17}, cervical¹⁶, prostate^{16,18}, colorectal^{16,18}, sleep apnea¹⁹.

Aim of this study was to assess the association of elevated body mass index and hypertension with general mortality in the Croatian adult population cohort from Croatian Adult Health Cohort Study (CroHort).

Sample and Methods

Sample

Respondents who participated in Croatian Adult Health Survey 2003 (9,070 respondents) were contacted in 2008 in order to participate in the follow-up Croatian Adult Health Cohort Study (CroHort). They were 18 years or older at the time of the first survey. The contact was established with 6,682 persons (73.7% of the original sample), out of which 3,229 agreed to participate in follow-up survey (35.6%), while 3,453 persons refused (38.1%). For 808 persons (8.9% of the original sample) information was obtained that they had died. Contact with remaining 1,580 persons (17.4%) was not established, meaning that there was no information available whether they were alive or not in 2008. Further details on this sample were described elsewhere²⁰.

Only persons for which information whether they were alive in 2008 was available, were included in the analysis encompassing total of 7,490 respondents (82.6% of the original sample), out of which 6,682 were alive in 2008 and 808 were dead.

Variables

Public health nurses measured blood pressure in respondents' homes, while weight and height used to calculate body mass index (BMI) were self-reported by respondents and recorded on a questionnaire by public health nurses. Blood pressure was measured twice during single visit using a mercury sphygmomanometer, with respondents seated for 20 minutes between the measurements.

Respondents whose BMI was lower than 25 kg/m² were classified as normal weight, those whose BMI was 25 kg/m² or higher, but lower than 30 kg/m² were classified as overweight, while those whose BMI was 30 kg/m² or higher were classified as obese.

Respondents who stated that they use anti-hypertensive drugs or whose average blood pressure calculated from two measurements was equal or higher than 140 mmHg for systolic or equal or higher than 90 mmHg for diastolic blood pressure, were categorized as hypertensive.

Age of respondents used in analysis was from year 2003 and data about risk factors (BMI, hypertension) were also from 2003 survey.

Statistical analysis

Percentage of respondents who died in period 2003–2008 was calculated separately for men and women in three main age groups (18–34 years, 35–64 years, 65 years and older) and in two additional age subgroups (35–49 years, 50–64 years) for each BMI category: normal weight, overweight and obese as well as for both blood pressure states: normotensive and hypertensive.

The multiple logistic regression analysis (adjusted for age) was used to calculate OR for death in period 2003–2008 depending on BMI (enrolled as continuous variable) and presence of hypertension (enrolled as binary variable: normotensive/hypertensive), separately for men and women in each age group: 18–34 years, 35–49 years, 50–64 years, 65 years and older.

All confidence intervals (CI) were calculated with 95% probability levels.

Statistical analysis was performed using SPSS (version 14.01; License: Croatian National Institute of Public Health).

Results

Both among men and women no significant differences in percentage of respondents who died were recorded among normal weight, overweight and obese respondents in the age groups: 18–34 years and 35–64 years as well as in the age subgroups: 35–49 years and 50–64 years, with exception of overweight men who had significantly lower percentage of respondents who died compared to those with normal weight in the age group 35–64 years and with exception of obese women who had significantly lower percentage of respondents who died compared to overweight women in the age subgroup 35–49 years.

Among men aged 65 years and older, significantly lower percentage of obese men died compared to normal weight men, while no significant differences were recorded among women in that age group (Table 1).

Among men, no significant difference between normotensive and hypertensive respondents in percentage of those who died was recorded in any age group, while among women aged 35–64 years percentage of hypertensive respondents who died was significantly higher compared to normotensive (Table 2).

For men, multiple logistic regression analysis showed that being hypertensive was associated with increased OR for death, while higher BMI was associated with lower OR for death among respondents aged 65 years and older. In younger age groups, neither BMI nor hypertension were associated with significant difference in OR for death.

For women, logistic regression analysis showed that being hypertensive was associated with increased OR for death among respondents aged 50–64 years, while neither BMI nor hypertension were associated with significant difference in OR for death among other age groups (Table 3).

Discussion

The results showed that being hypertensive was associated with increased OR for death in man aged 65 years and older and women aged 50–64 years. A correlation between mortality and high blood pressure has been widely shown. Several studies have showed that the mortality rate was higher among hypertensive adults than non hy-

TABLE 1
NUMBER AND PERCENTAGE OF RESPONDENTS WHO DIED 2003–2008 ACCORDING TO THEIR WEIGHT IN 2003

Characteristic	Men			Women		
	Number	%	95% CI	Number	%	95% CI
Age 18–34						
normal weight (BMI < 25)	8	4.3	1.4–7.2	13	2.0	0.9–3.1
overweight (25 ≤ BMI < 30)	5	5.3	0.8–9.8	4	3.1	0.1–6.2
obese (BMI ≥ 30)	0	0.0		2	4.2	0.0–9.8
Age 35–64 total						
normal weight (BMI < 25)	35	10.5	7.2–13.8	53	5.5	4.0–6.9
overweight (25 ≤ BMI < 30)	33	5.4	3.6–7.1	60	6.3	4.8–7.8
obese (BMI ≥ 30)	22	6.4	3.8–9.0	31	5.0	3.3–6.7
Age 35–49						
normal weight (BMI < 25)	13	7.8	3.7–11.9	18	3.1	1.7–4.5
overweight (25 ≤ BMI < 30)	13	4.8	2.3–7.3	22	5.3	3.1–7.5
obese (BMI ≥ 30)	8	5.7	1.9–9.6	3	1.4	0.0–3.0
Age 50–64						
normal weight (BMI < 25)	22	13.1	8.0–18.2	35	9.0	6.1–11.8
overweight (25 ≤ BMI < 30)	20	5.8	3.3–8.3	38	7.1	4.9–9.2
obese (BMI ≥ 30)	14	6.9	3.4–10.4	28	6.8	4.4–9.3
Age 65 and older						
normal weight (BMI < 25)	79	31.2	25.5–36.9	123	24.6	20.9–28.4
overweight (25 ≤ BMI < 30)	91	23.6	19.3–27.8	129	18.5	15.6–21.4
obese (BMI ≥ 30)	25	15.4	9.9–21.0	93	18.4	15.0–21.8

TABLE 2
NUMBER AND PERCENTAGE OF RESPONDENTS WHO DIED 2003–2008 ACCORDING TO THEIR BLOOD PRESSURE IN 2003

Characteristic	Men			Women		
	Number	%	95% CI	Number	%	95% CI
Age 18–34						
normotensive	9	3.6	1.3–5.9	19	2.5	1.4–3.7
hypertensive	4	6.3	0.3–12.4	0	0.0	
Age 35–64 total						
normotensive	45	7.4	5.3–9.5	60	4.3	3.3–5.4
hypertensive	45	6.5	4.7–8.4	85	7.3	5.8–8.8
Age 35–49						
normotensive	23	6.9	4.2–9.6	32	3.7	2.4–4.9
hypertensive	11	4.5	1.9–7.1	12	3.5	1.6–5.5
Age 50–64						
normotensive	22	8.1	4.8–11.3	28	5.4	3.5–7.4
hypertensive	34	7.6	5.2–10.1	73	8.8	6.9–10.8
Age 65 and older						
normotensive	38	19.0	13.6–24.4	60	17.9	13.8–22.0
hypertensive	157	26.0	22.5–29.5	286	20.9	18.8–23.1

pertensive adults^{21–23}. The public health significance of hypertension results from the finding that it represents an independent risk factor for cardiovascular diseases. Progression of the hypertension is associated with vascular abnormalities that damage the heart, kidneys, brain

and other organs, and leads to premature morbidity and death^{24–28}.

Compared to normotensive individuals, people with an elevated blood pressure have an increased tendency to have other risk factors for CVD (i.e. diabetes, dyslipidemia)²⁹.

TABLE 3
ASSOCIATION OF BMI AND HYPERTENSION WITH RISK OF
DEATH IN PERIOD 2003–2008

Characteristic	Men		Women	
	OR*	95% CI	OR*	95% CI
Age 18–34				
BMI	0.93	0.79–1.10	1.09	0.98–1.22
hypertension	2.08	0.59–7.32	0.00	
Age 35–49				
BMI	0.97	0.90–1.06	1.02	0.96–1.10
hypertension	0.67	0.31–1.45	0.87	0.42–1.79
Age 50–64				
BMI	0.93	0.87–1.00	0.97	0.93–1.02
hypertension	1.10	0.62–1.97	1.65	1.03–2.65
Age 65 and older				
BMI	0.93	0.89–0.98	0.99	0.96–1.01
hypertension	1.53	1.01–2.30	1.25	0.91–1.72

* adjusted for age

Systolic and diastolic blood pressure tends to rise with age and prevalence of hypertension increases after the age of 50 years. Some population studies have showed positive relationship between blood pressure and mortality at the age of 60–69 years, less in age group 75 years and older, but negative relationship in older than 85 years. This negative relationship in very elderly is probably caused by terminal phase of cardiac, respiratory and neoplastic disease and consequent low blood pressure³⁰.

Analysis showed that in the age group 35–64 years significantly lower percentage of respondents died among overweight men compared to those with normal weight and in the age subgroup 35–49 years significantly lower percentage of respondents died among obese women compared to overweight women. Among men aged 65 years and older, higher BMI was associated with lower OR for death.

In several studies overweight and obesity are associated with increased all-cause mortality and showed an optimal BMI of 22.5 to 25.0 in analyses of all study participants^{31–33}.

Obesity has been clearly identified as a risk for premature death^{34–36}, but the most studies recorded variability in the association of overweight and mortality. Flegal et al., Heather M. Orpana et al. and other authors found that underweight and obesity (particularly higher levels of obesity) were clear risk factors for mortality, but overweight was associated with a significantly decreased risk of death^{37–39}.

Association between elevated body weight and excess deaths is not necessarily causal. Many factors influence mortality risk in addition to higher BMI. The reason for variability among studies assessing the association of BMI and mortality was that they had a different length of follow-up and inadequate control for confounder that

might mask association of BMI with mortality⁴⁰. This study has a short follow-up and it is possible that the pattern of results could change during a longer period of follow-up. The relation between BMI and all causes death differed according to smoking status, pre-existing chronic diseases and weight loss due to disease which may or may not be recognizable clinically^{38,41–44}.

Due to the concern that pre-existing disease could influence the association between BMI and mortality, our analyses were also adjusted for hypertension, but not for other chronic diseases.

Smoking is an important confounder because people who smoke generally have a lower body mass index than non-smokers^{41,45}.

People who smoke have a higher risk of death due to cardiovascular diseases (heart attack and stroke, peripheral vascular disease), chronic obstructive pulmonary diseases or cancer and low weight or weight loss may be in relationships with ill health. When assessing the relationship between body weight and mortality, it is important to taking smoking status and pre-existing disease into account^{46,47}.

In the general population, obesity is associated with increased risk of adverse outcomes. Studies of patients with chronic disease suggest that overweight and obese patients may paradoxically have lower risk of death compared with those with normal weight. Pre-existing illness could lead to lower BMI and earlier mortality^{43,47,48}.

In most reviewed studies, optimal BMI was associated with the lowest mortality increase with age and higher weight is not often associated with poor health^{49–53}.

The results of this research support the obesity paradox presence in the elderly. They showed that being obese is a protective factor for death in men aged 65 years and older. Elderly with low BMI or weight loss may have chronic disease, co-morbidity and unrecognized illness which can be associated with higher mortality rates⁵⁴. In most populations, the body-mass index is correlated with more direct measures of body fat, but in elderly it may be a less useful indicator of adiposity. BMI may be relatively low among older people who have little muscle mass compared to adipose tissue and who tend to have a shift of fat from peripheral to central sites without increase in body-mass index⁵⁵.

Many factors influence body weight and additional research to confirm association between BMI and mortality in different age groups would be of interest.

Excess body weight and hypertension are strong determinants as well as modifiable risk factors for all-cause mortality, and as such carry the potential for primary prevention. Despite variability in association of overweight and mortality among different age groups, a review of the literature provides much support for public health messages that advocate the benefit of change to healthier lifestyle regardless of age, in order to prevent excess body weight and hypertension as well as associated premature mortality.

Acknowledgements

This article was prepared as a part of scientific project »Regionalism of cardiovascular behavioural risk factors –

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POVEZANOST POVEĆANOG INDEKSA TJELESNE MASE I HIPERTENZIJE S MORTALITETOM U HRVATSKOJ KOHORTNOJ STUDIJI KARDIOVASKULARNOG ZDRAVLJA 2003/08

SAŽETAK

Čilj ovog rada bio je istražiti povezanost povećanog indeksa tjelesne mase (ITM) i hipertenzije s općim mortalitetom u kohorti ispitanika Hrvatske kohortne studije kardiovaskularnog zdravlja 2003/08. Rizik od smrti prema kategorijama

indeksa tjelesne mase i krvnog tlaka, u razdoblju 2003–08. godine, izračunat je za 7,490 ispitanika, od kojih je u 2008. godini bilo 6,682 živih i 808 mrtvih. Kod muškaraca u dobi od 65 i više godina, povećan ITM bio je povezan s manjim rizikom od smrti u skladu s paradoksom debljine zabilježenim u nekim prethodnim studijama koje su pokazale sa stariji muškarci višim ITM imaju manji rizik od smrti. Kod žena u dobi 50–64 godine, prisutnost hipertenzije bila je povezana s povećanim rizikom od smrti u usporedbi s ispitanicama s normalnim krvnim tlakom. Usprkos paradoksu debljine koji traži daljnja istraživanja, potrebno je poticati aktivnosti u primarnoj i sekundarnoj prevenciji prekomjerne tjelesne težine i hipertenzije kao sredstvima za sprječavanje prerane smrtnosti u hrvatskom stanovništvu.