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# Biological fitness at middle-age is reduced in both very lean and obese males

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*ABSTRACT* It is a well-documented fact that the relationship between body weight and premature mortality is not linear. Very often it is described as J- or U-shaped. There is solid evidence for strong relationships in adults between obesity (as defined by high values of the body mass index, BMI), mortality, hypertension, cardiovascular disease, stroke, and non-insulin dependent diabetes. However, the opposite end of the BMI distribution (leanness) remains an area of controversy among European populations. Many authors have indicated that leanness elevates the risk of premature mortality, although to a lesser degree than obesity. The aim of this study is to examine whether abnormalities in blood pressure and lung function occur more frequently at both extremes of the BMI distribution than in its middle range.

KEY WORDS blood pressure, respiratory indices, J-shaped distribution, BMI

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## Introduction

Many studies have shown a relationship between body weight and premature mortality. This association has been described as being J-shaped, U-shaped, inverse, positive, or absent [Report of a WHO Expert Committee 1995, KUSHNER 1993, MÅNSSON *et al.* 1996, ALLISON *et al.* 1997]. There is solid evidence for strong relationships in adults between obesity (as defined by high values of the body mass index, BMI), mortality, hypertension, cardiovascular disease, stroke, and non-insulin dependent diabetes [JOOS *et al.* 1984, DONAHUE *et al.* 1987]. For example, in obese patients from Düsseldorf, Germany, the mortality risk ratio for men with a BMI between 36 and 40 kg/m<sup>2</sup> is 1.50 and for men with BMI equal to or higher than 40 kg/m<sup>2</sup> it is 2.10, relative to a reference group of a BMI between 25 and 32 [BENDER *et al.* 1998]. Also, physiological factors increasing the risk of mortality, such as high levels of cholesterol and glucose, are more frequent among obese than normal weight adults [OSTLUND *et al.* 1990].

However, the opposite end of the BMI distribution remains an area of controversy among European populations. Many authors have indicated that leanness elevates the risk of premature mortality, although to a lesser degree than obesity [RHOADS and KAGAN 1983, DORN et al. 1997, THOROGOOD et al. 2003]. The observation that individuals with the lowest relative weights die earlier than individuals with intermediate levels of the BMI, has been questioned. Several authors have expressed doubt about such a relationship and have treated it as an artifact. They have pointed out that cigarette smoking or subclinical diseases could confound the relationship between body weight and mortality because of their direct link with low body weight [MANSON et al. 1995, ALLISON et al. 1997].

The aim of this study is to examine whether abnormalities in blood pressure and lung function occur more frequently at both extremes of the BMI distribution than in its middle range.

### Materials and methods

Between 1982 and 1992, anthropological and medical examinations of 1073 males, 40-50 year-old inhabitants of Wrocław, Poland, were carried out at the Institute of Anthropology, Polish Academy of Sciences, and the Lower Silesian Medical Center, Wrocław. Three social groups of men: professionals, skilled, and unskilled workers were investigated. For the purpose of the present analysis, only men who smoked were chosen.

The following four health-related functional traits were measured: systolic blood pressure SBP (mmHg); diastolic blood pressure DBP (mm Hg); relative vital capacity RVC (ratio of vital capacity to body height); and forced expiratory volume in 1 sec. FEV (ml). Systolic and diastolic blood pressure (SBP, DBP) was measured twice in the sitting position using an MPC-350 apparatus and the two readings averaged for data analyses. Respiratory characteristics were collected using a Collins apparatus, according to standard procedures. Body height and body weight were measured and the body mass index (BMI) was calculated.

The sample was divided into three groups, according to BMI values: (1) lean (BMI equal to or under 20, N = 97); (2) medium (BMI between 20 and 30, N = 867; and (3) obese (BMI over 30, N = 109). The WHO thresholds [Report of a WHO Expert Committee, 1995] were not used in this paper: Among 40-50-year old men, there were not enough lean individuals with a BMI under 18; hence, this threshold was raised to 20. Individuals with grade I of overweight (BMI 25-30) were included in the group with medium weight. It allowed inclusion of the third group only persons with grades II and III of overweight.

Four separate one-way analyses of variance were used to compare healthrelated functional traits of men across the BMI groups. To assess the shape of the analyzed relationship, the values for given characteristics in each group of the BMI were compared with the threshold values (25th or 75th centile) based on an optimal group. The optimal group was a 25-29-age group of college educated men [ROGUCKA and WELON 1996]. From the biological and medical points of view, 25-29 year-old males have reached full physiological maturity and their health-related characteristics have achieved their most favorable values. Next, for each group of men in respect of the characteristics increasing with age (blood pressure), the proportions of men whose values were above the 75th percentile were calculated. For characteristics decreasing with age (respiratory indices), the proportions of men whose values were below the 25th percentile were calculated. To test the differences between the two proportions, a chi-square test was done.

## Analysis

The results of the four separate oneway analyses of variance are summarized in Table 1. There was a significant correlation between the BMI and three out of four analyzed functional healthrelated traits (systolic blood pressure, diastolic blood pressure, and forced expiratory volume in 1 sec.). Relative vital capacity did not show any significant correlation with the body mass index.

The values of the 25th or the 75th centile (depending on a negative or a positive linear trend with age) of four functional health-related traits, proportions of lean, medium and obese men (according to the BMI values), and the level of significance of differences in frequencies between the analyzed groups of men are presented in Table 2. The percentage of males with systolic blood pressure exceeding the 75th centile of that trait for young well-educated males (132.2 mm Hg) was only 38.5% among the subjects

 Table 1. Results of four separate one-way analyses of variance for health-related functional traits of men across the BMI groups

Dependent variable	Main effect		
	F	p	
Systolic blood pressure (SBP)	14.84	0.000	
Diastolic blood pressure (DBP)	13.65	0.000	
Forced expiratory volume (FEV) in 1 sec.	4.95	0.007	
Relative vital capacity (RVC)	1.68	0.186	

**Table 2**. Proportion of men (aged 40-50) whose values were above the 75th percentile (for SBP and DBP) and below the 25th percentile (for both respiratory indices), based on an optimal group (aged 25-29)

	Threshold 25/75 centile	(1) Lean	(2) Medium	(3) Obese	Significance between pairs
SBP	132.2	50.8 %	38.5 %	61.7 %	(1-2) * (2-3)***
DBP	89.8	36.9 %	27.1 %	48.1 %	(1-2) * (2-3)***
RVC	24.8	70.0 %	66.5 %	75.6 %	
FEV in 1 sec.	3687	84.4 %	69.1 %	76.9 %	(1-2)***

Level of significance: \* p < 0.05; \*\*\* p < 0.001

with medium values of the BMI, but almost 51 % among the very lean and almost 62% among the obese ones. A similar pattern occurred when diastolic blood pressure was analyzed. As many as 48.1% of obese men and 36.9% of lean men had diastolic blood pressure higher than 89.9 mmHg, whereas only 27.1% of men with medium weight exceeded this value. The percentage of men with relative vital capacity who did not reach the 25th centile for that trait for young well-educated males was 66.5% among the subjects with medium values of the BMI, whereas it was as much as 70% among the very lean and 75.6% among the obese ones. Similarly, the percentage of men with forced expiratory volume in 1 sec. who did not reach 3687 ml was 69.1% among the subjects with medium weight, whereas it reached as much as 84.4% among the very lean and 76.9% among the obese ones. The non-linear relationship between the body mass index and functional characteristics is presented graphically in Figures 1-4.

## Discussion

A J-shaped relationship between the BMI and functional characteristics is observed in this study. Among males aged 40-50 and economically active inhabitants of Wrocław, those with BMI between 20 and 30 had the bestfunctioning circulatory and respiratory systems. Extreme fatness and extreme leanness were both associated with an impaired function of these systems. Men with BMI equal to or under 20, and men with BMI over 30 achieved higher values of systolic and diastolic blood pressure and lower values of forced expiratory volume in 1 sec. Only relative vital capacity did not exhibit such a relationship.

A number of empirical studies have shown that a similar pattern occurred when the association between mortality ratios and BMI values was analyzed. DORN et al. [1997] investigated the long-term relation between the body mass index and mortality among men and women aged 20-96 in the Buffalo Blood Pressure Study. A significant linear association was found between the BMI and all-cause mortality in men under 65 years old, but not in women. In men aged 65 and older, the relation was quadratic in form (p = 0.02), with the lowest risks appearing in a BMI range of 23-27. RHOADS and KAGAN [1983] found that among 45-68 year-old Honolulu Japanese men that mortality was highest in both the leanest and fattest men and lowest in the fourth quintile of the body mass index. An excess of deaths in the top quintile was caused primarily by coronary heart disease (CHD), whereas in the bottom two quintiles the excess deaths were due to cancer and "other" causes. In addition, in an unusually slim group of non-meat eaters from the United Kingdom, participants with a BMI below 18 had an increased all cause death ratio and increased ratios of death from circulatory and respiratory diseases [THOROGOOD et al. 2003].

Several authors have suggested that the relation described above is an artifact and that the increased risk of mortality at low BMI values is not real. In this context two confounding variables were mentioned: effects of the smoking habit, and pre-existing diseases. It is a well-known fact that cigarette smoking

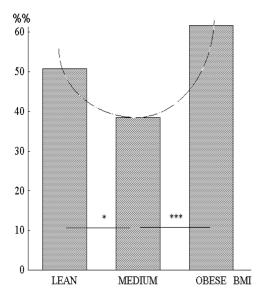


Fig 1. The proportion of males with systolic blood pressure exceeding the 75th centile, based on optimal group.

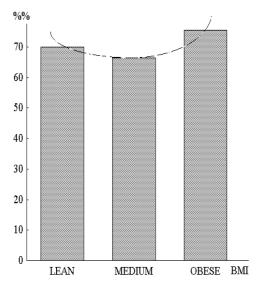


Fig. 3. The proportion of males with relative vital capacity below the 25th centile, based on optimal group.

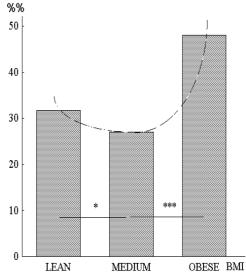


Fig. 2. The proportion of males with diastolic blood pressure exceeding the 75th centile, based on optimal group.

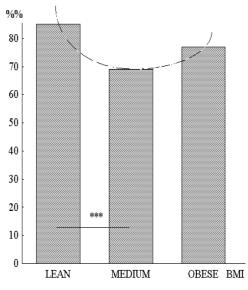


Fig. 4. The proportion of males with forced expiratory volume in 1 sec. below the 25th centile, based on optimal group.

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is responsible for both low body weight, due to suppression of appetite and increasing body metabolic rates [HICKEY and MULCAHY 1973, HOFSTETTER et al. 1986], and for higher mortality rates [WYNDER and HOFFMANN 1967, DOLL and PETO 1981]. MANSON et al. [1995] found that a J-shaped association between the BMI and overall mortality existed among women aged 30-55. However, when women who had never smoked were examined separately, the J-shaped association disappeared. It remained only among formerly and currently smoking women. Furthermore, in the case of pre-existing diseases, when loss of weight is often a symptom of an early death, the exclusion from longterm analysis of subjects who had died during the first several years caused the disappearance of the J- or U-shaped association.

MANSON et al. [1995] claimed that excess risks of mortality associated with leanness in their study were artifactual and were therefore eliminated after accounting for cigarette smoking and subclinical disease. However, many studies controlling these confounders [RISSANEN et al. 1989, FOLSOM et al. 1993, SINGH et al. 1999, MEYER et al. 2002] still show the J- or U-shaped relation. SINGH et al. [1999] found a J-shaped relation between body weight and mortality among older women who had not undergone postmenopausal hormone replacement. Next, MEYER et al. [2002] suggested that a J-shaped association between the BMI and total mortality existed even when smoking habits and physical activity were analyzed. MÅNSSON et al. [1996] examined adult males from Malmö, Sweden, and found that among under- and overweight,

especially the obese, a high proportion of males were on disability pensions. After adjustment for smoking, there was still a J-shaped relation between the body mass index and the incidence of disability pensions. The relative risk (with a group of normal BMI as reference) among underweight men was 1.9, for the overweight subjects -1.3 and for the obese -2.8, all differences being significant. After adjustment for hypertension, these results persisted, although slightly weakened. The high pension rate among the obese men was mainly a result of diseases of the circulatory and musculoskeletal systems (diseases of muscles and connective tissue). Among lean men, the disability pension was granted primarily because of alcohol dependence and diseases of the musculoskeletal system (collagenoses, including rheumatoid arthritis and diseases of muscles and connective tissue).

ALLISON et al. [1997] presented an additional hypothesis to explain the Uor J-shaped relation between the BMI and mortality. The elevated risk of mortality at the low end of the BMI distribution may be due to a measurement problem, namely the use of the BMI as a measure of "adiposity". These authors suggested that this relation might result from the fact that the BMI is composed of separate components, mainly fat mass (FM) and fat-free mass (FFM), which have opposite effects on health and longevity. Increasing amounts of FM pose a threat to health [STEVENS and MARQUIS 1996], while increasing amounts of FFM enhance it [SHEPHARD 1994]. At the same time BERGMAN and ROGUCKA [1998] found that for men aged 40-50 the correlation between the BMI and other mea-

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surements of fatness (e.g., the sum of skinfolds, impedance) was higher for lean persons (BMI equal to or under 24) than for obese ones (BMI equal to or over 27).

In conclusion, the BMI is a good indicator of the risk of morbidity and mortality, at least in middle-aged adults. In many populations the J- or U-shaped relationship between relative weight and all-cause mortality has been reported. It is assumed that causes of death associated with a low BMI are tuberculosis, obstructive lung disease, and cancer of the lung and stomach; whereas those associated with a high BMI are cerebrovascular disease, cardiovascular disease, diabetes and, in men, colon cancer [Report of a WHO Expert Committee, 1995]. Few explanations for such a pattern are offered (from artifact to behavioral reasons), but none explains the problem satisfactorily. In this study, among males aged 40-50, both obesity and extreme leanness were associated with an impaired function of the circulatory and respiratory systems. It reflects the J- or U-shaped relationship between body weight and mortality.

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#### Streszczenie

Wiele badań nad związkiem przedwczesnej umieralności i masy ciała wskazuje na istnienie J- lub U-kształtnej zależności. Podczas gdy zwiększona umieralność oraz zachorowalność wśród osób otyłych jest dobrze udokumentowana, to pytanie czy zależność ta dotyczy również osób bardzo szczupłych, wciąż budzi wiele kontrowersji. Wielu autorów dowodzi, że nadmierna szczupłość ciała również podwyższa ryzyko przedwczesnej śmierci, choć w mniejszym stopniu niż otyłość. Celem tego badania jest odpowiedź na pytanie, czy nieprawidłowości w funkcjonowaniu układu krwionośnego oraz układu oddechowego, pojawiają się częściej na obu końcach rozkładu względnej masy ciała (BMI), a więc wśród osób otyłych i bardzo szczupłych, niż wśród osób o średniej masie ciała.

Analizę oparto na danych pomiarowych 1073 mężczyzn, 40-50-letnich, aktywnych zawodowo mieszkańców Wrocławia palących tytoń. Do oceny stanu układu krążenia użyto skurczowego i rozkurczowego ciśnienia krwi; do oceny stanu układu oddechowego wykorzystano względną pojemność życiową płuc (pojemność życiowa płuc/wysokość ciała) oraz jednosekundową pojemność wydechową płuc. Badanych mężczyzn podzielono na trzy grupy: szczupłych (BMI  $\leq 20$ , N = 97); o średniej masie ciała ( $20 < BMI \leq 30$ , N = 867) oraz otyłych (BMI > 30, N = 109). W celu porównania wartości poszczególnych analizowanych zmiennych w trzech wyróżnionych ze względu na BMI grupach mężczyzn zastosowano cztery oddzielne jednoczynnikowe analizy wariancji. Wyniki poszczególnych ANOVA zawiera Tab. 1. Trzy spośród czterech analizowanych zmiennych wykazały istotną zależność od BMI (skurczowe ciśnienie krwi: F = 14,84; p = 0,000; rozkurczowe ciśnienie krwi: F = 13,65; p = 0,000; oraz jednosekundowa pojemność wydechowa płuc: F = 4,95; p = 0,007). Względna pojemność życiowa płuc nie wykazała istotnej zależności od BMI.

Następnie dla każdego analizowanego parametru układu krążenia i oddechowego, obliczona została wartość graniczna, od której możemy mówić o nieprawidłowości. Dla cech, których wartości wzrastają z wiekiem (ciśnienie krwi) jest to 75 centyl, natomiast dla cech, których wartości obniżają się wraz z wiekiem (parametry oddechowe) – 25 centyl, wyliczony z rozkładu danej cechy w populacji zdrowych, 25-29 letnich mężczyzn z wyższym wykształceniem. W każdej z trzech grup mężczyzn, obliczono procentowy udział osób, przekraczających (lub nie osiągających) określone wartości graniczne analizowanych parametrów. Poszczególne wartości graniczne oraz częstości mężczyzn pozostających poza tymi granicami zawiera Tab. 2. Wśród mężczyzn ze średnią masą ciała tylko 38,5% posiadało ciśnienie skurczowe krwi powyżej 132,2 mmHg, podczas gdy wśród otyłych, takich osób było aż 61,7%, a wśród bardzo szczupłych 50,8%. Podobnie było w przypadku ciśnienia rozkurczowego krwi: 48,1% otyłych i 36,9% szczupłych badanych miało ciśnienie rozkurczowe krwi powyżej 89,9 mmHg, podczas gdy tylko 27,1% mężczyzn ze średnią masą ciała osiągnęło wartość graniczną tego parametru. Podobna zależność widoczna była w przypadku parametrów oddechowych. Wśród mężczyzn z normalną masą ciała 66,5% miało relatywną pojemność życiową płuc poniżej wartości 24,8, podczas gdy wśród otyłych 75,6%, a wśród szczupłych – 70,0% nie osiągnęło wartości progowej dla tej cechy (różnice nieistotne statystycznie – patrz Tab. 2). Ponadto 69,1% mężczyzn ze średnią masą ciała miało jednosekundową pojemność wydechową płuc poniżej wyznaczonego poziomu progowego wynoszącego 3687 ml, podczas gdy wśród otyłych było 76,9%, a wśród szczupłych – aż 84,4% takich osób. Rys. 1-4 przedstawiają powyższe zależności w sposób graficzny.

Przedstawione wyniki jednoznacznie wskazują na istnienie J-kształtnej zależności pomiędzy masą ciała a parametrami układu krążenia i układu oddechowego, gdzie zarówno otyłość (BMI > 30) jak i nadmierna szczupłość ciała (BMI  $\leq$  20) upośledza funkcjonowanie organizmu mężczyzn w wieku 40-50 lat. Odzwierciedleniem tego jest taka sama zależność umieralności od względnej masy ciała. Powstało kilka prac, których autorzy starają się wyjaśnić tę zależność (od artefaktu do przyczyn związanych ze stylem życia), jednakże żadna z nich nie tłumaczy ich w pełni satysfakcjonująco.