

conditions, MHO and fit obese, have not yet been adequately investigated, thereby leaving a void that requires further exploration.

In the recent paper by Chang et al. (8), coronary artery calcium (CAC) scores were assessed among Korean subjects; they had a very low prevalence of obesity and CAC compared with people in the United States and India but had higher CAC scores, even among those with MHO. Although the investigators tried to assess the performance of regular exercise, there was no detailed assessment of physical activity or CRF. Future studies should combine the important components of metabolic health and CRF to determine the impact of adiposity on cardiovascular disease. The need for a healthy metabolic profile and CRF cannot be understated and should be promoted, irrespective of body mass index or CAC score.

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<http://dx.doi.org/10.1016/j.jacc.2014.05.061>

Please note: Dr. Lavie is the author of *The Obesity Paradox*. All other authors have reported that they have no relationships relevant to the contents of this paper to disclose.

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REPLY: How to Determine a Metabolically Healthy Body Composition in Cardiovascular Disease



We thank Dr. Babu and colleagues and Dr. Scherbakov and colleagues for their interest in our paper (1) and for addressing important points in the interpretation of our results. Several of the arguments of Dr. Scherbakov and colleagues address the issue of categorization of parameters that have an underlying continuous distribution, such as coronary artery calcium (CAC) scores or body mass index (BMI). The actual cutoffs used for these parameters are arbitrary and established for clinical convenience but do not reflect changes in the underlying biological processes. With regard to CAC, cutoffs have been developed for risk stratification; however, CAC scores have a direct continuous relationship to total coronary plaque burden in histology (2), severity of coronary artery disease (3), and risk of coronary artery disease, including comparisons of very low CAC scores with a score of 0 (4,5). In our analysis, we used a cutoff of 80 Agatston units (6,7) in multinomial logistic regression models because of the low prevalence of elevated CAC scores; only 13 subjects (0.1%) had a CAC score >400, and 80 Agatston units corresponded approximately to the 90th percentile of participants with any detectable CAC. These results were consistent with alternative analytical models presented in the paper, including using CAC scores as a continuous outcome. Regardless of the analytical methods, metabolically healthy obese participants had a higher prevalence of subclinical coronary atherosclerosis compared with metabolically healthy participants with normal weight, and this association persisted after adjustment for potential confounders, including sex, cigarette smoking, and alcohol consumption.

With regard to BMI, there is an extensive body of evidence that, for the same level of BMI, Asian subjects have higher total body fat (8) and a higher risk of cardiometabolic disease (9) compared with white subjects. Thus, a World Health Organization consultation proposed Asian Pacific-specific BMI criteria (10), which have gained widespread acceptance. Irrespective of the cutoffs and the characteristics of the study populations, the mechanisms linking higher BMI to subclinical atherosclerosis are likely to be similar across populations.

Both Dr. Babu and colleagues and Dr. Scherbakov and colleagues discuss the obesity paradox and indicate that BMI has a U-shaped relationship with mortality. The obesity paradox is a complex phenomenon and affects primarily studies of subjects older than

50 years of age or with established comorbidities (11-15), raising concerns about selection, survival, treatment, and confounder biases (14). The relevance of the obesity paradox to our study population, composed of relatively young asymptomatic Korean men and women, is uncertain. There is, however, an extensive body of evidence indicating that BMI is linearly associated with the development of metabolic abnormalities such as insulin resistance, type 2 diabetes mellitus, and hypertension (14,16) and with cardiovascular mortality and incidence in the Asian population (17,18).

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