




## Reply to Flegal

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We thank Dr Flegal for her interest in our recent study. We understand her concerns about self-reported anthropometric data and acknowledge the limitation that weight was not measured in all 10 cohorts, as we noted in the Discussion section of the article. Reassuringly, validation studies have shown that the absolute difference between reported and measured weight is relatively small. In the anthropometric validation study from the Million Women Study cohort, Wright et al. (1) documented a mean (SD) reported weight of 68.4 (11.9) kg and measured weight of 69.8 (12.6) kg among 40 000 women with both measures at the same time point. The calculated regression dilution ratio for weight in that study was 1.02, and the authors concluded that “reporting errors are likely to generate very little bias in estimates of associations with disease outcomes” (1). In her letter, Dr Flegal points out a small but statistically significant difference in the hazard ratios for self-reported and measured data from one subset of studies in the Global BMI Mortality Collaboration Cohort (2). However, the difference in the results obtained using self-reported and measured body mass index when all 239 studies were included was not statistically significant. Furthermore, the same conclusions were drawn regardless of whether the weight was self-reported or measured, a modest increased risk of death for overweight and a greater risk for obesity.

In reference to Dr Flegal’s concern about our categorical variable, we also conducted a spline regression analysis of sustained weight change as a continuous variable (analysis restricted to participants with sustained weight loss, sustained weight gain, or stable weight). Using stepwise selection, none of the spline variables remained in the model, providing further evidence of a linear dose response with increasing amounts of sustained weight loss.

Most important, in a sensitivity analysis stratifying our main analysis on anthropometric ascertainment method, we observed a similar inverse association in both groups (Supplementary Table 8 in the original article). In fact, the

hazard ratios were stronger in the measured compared with self-report group. The confidence intervals were also a bit wider, as would be expected based on the relative sample size of the two groups. Our results were robust in every sensitivity analysis; regardless of the method of weight ascertainment, sustained weight loss in women aged 50 years and older was inversely associated with breast cancer risk in a dose-related fashion. These results, taken together with the other scientific evidence documenting the relationships between weight change and sex steroid hormone levels and between hormone levels and risk of breast cancer, are encouraging for breast cancer prevention (3–6).

### Notes

We declare no competing interests. Lauren R. Teras, Alpa V. Patel, and Stephanie A. Smith-Warner for the Pooling Project of Prospective Studies of Diet and Cancer (lteras@cancer.org).

### References

1. Wright FL, Green J, Reeves G, Beral V, Cairnes BJ. Validity over time of self-reported anthropometric variables during follow-up of a large cohort of UK women. *BMC Med Res Methodol*. 2015;15(1):81.
2. Di Angelantonio E, Bhupathiraju SN, Wormser D, et al. Body-mass index and all-cause mortality: individual-participant-data meta-analysis of 239 prospective studies in four continents. *Lancet*. 2016;388(10046):776–786.
3. Key TJ, Appleby PN, Reeves GK, et al. Steroid hormone measurements from different types of assays in relation to body mass index and breast cancer risk in postmenopausal women: reanalysis of eighteen prospective studies. *Steroids*. 2015;99(Pt A):49–55.
4. de Waard F, Poortman J, de Pedro-Alvarez Ferrero M, Baanders-van Halewijn EA. Weight reduction and oestrogen excretion in obese post-menopausal women. *Maturitas*. 1982;4(2):155–162.
5. O’Dea JP, Wieland RG, Hallberg MC, Llerena LA, Zorn EM, Genuth SM. Effect of dietary weight loss on sex steroid binding sex steroids, and gonadotropins in obese postmenopausal women. *J Lab Clin Med*. 1979;93(6):1004–1008.
6. van Gemert WA, Schuit AJ, van der Palen J, et al. Effect of weight loss, with or without exercise, on body composition and sex hormones in postmenopausal women: the SHAPE-2 trial. *Breast Cancer Res*. 2015;17(1):1.

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