receptor only. However, the actual role of androgen receptor in breast cancer remains unclear. Considering the relatively consistent finding of hyperandrogenism in breast cancer, the impact of androgen on breast cancer warrants further investigation.

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## References

- Cauley JA, Lucas FL, Kuller LH, Stone K, Browner W, Cummings SR. Elevated serum estradiol and testosterone concentrations are associated with a high risk for breast cancer. Ann Intern Med. 1999;130:270-7.
- Dorgan JF, Longcope C, Stephenson HE, Falk RT, Miller R, Franz C, et al. Relation of prediagnostic serum estrogen and androgen levels to breast cancer risk. Cancer Epidemiol Biomarkers Prev. 1996;5:533-9.
- Berrino F, Muti P, Micheli A, Bolelli G, Krogh V, Sciajno R, et al. Serum sex hormone levels after menopause and subsequent breast cancer. J Natl Cancer Inst. 1996;88:291-6.
- Hankinson SE, Willett WC, Manson JE, Colditz GA, Hunter DJ, Spiegelman D, et al. Plasma sex steroid hormone levels and risk of breast cancer in postmenopausal women. J Natl Cancer Inst. 1998;90:1292-9.
- Yu H, Bharaj BS, Giai M, Diamandis EP. CAG repeats of the androgen receptor gene in breast cancer and its association with survival [Abstract]. AACR Proceedings. 1999;40:380.

## **Risk for Breast Cancer**

To the Editor: Cauley and colleagues (1) reported that breast cancer risk was elevated not only in white postmenopausal women with high estrogen levels but also in those with high androgen levels. The association of breast cancer risk with androgen has been seen in many epidemiologic studies (2–4), but the mechanism by which androgens contribute to the cause of breast cancer is poorly understood. The relation between androgen and breast cancer is believed to be confounded by the strong correlation between androgen and estrogen because androgen is a precursor of estrogen in its metabolic pathway. Cauley and coworkers found that free testosterone was linked to breast cancer risk after adjustment for estrogens, suggesting that the association of androgen with breast cancer is independent of estrogen.

In a recent study by our group (5), we also found evidence supporting the notion that androgen may play a role in breast cancer. In breast cancer tissue, we analyzed DNA sequence from exon 1 of the androgen receptor gene that contained the variable length of CAG repeat known to inversely affect transcription according to its length. Our study showed that compared with longer CAG repeats, shorter CAG repeats were associated with aggressive disease and poor survival. This finding suggests that enhanced transcriptional activity of the androgen receptor gene might promote breast cancer progression.

Androgen receptor is expressed more frequently in breast cancer tissue than the estrogen or progesterone receptor is. This indicates that a subset of breast tumors may express androgen