



Obesity and Breast Cancer: The Role of Dysregulated Estrogen Metabolism (SpringerBriefs in Cancer Research)

Kristy A. Brown, Evan R. Simpson

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Obesity is a risk factor for breast cancer in older women. A number of adipose-derived and obesity-related factors have been shown to affect tumour cell growth. These include adipokines, insulin, IGF-1 and oestrogens. The majority of obesity-related postmenopausal breast cancers are oestrogen-dependent. Since the ovaries no longer produce oestrogens after menopause, and that circulating levels are negligible, it is evident that it is the oestrogens produced locally within the breast adipose that are responsible for the increased growth of breast cancer cells. Aromatase is the enzyme that converts androgens into oestrogens and its regulation is dependent on the activity of a number of tissue-specific promoters. Targeting oestrogen biosynthesis in obesity may be useful for the prevention of breast cancer. Aromatase inhibitors are efficacious at treating postmenopausal breast cancer and recent studies suggest that they may also be useful in the prevention setting. However, these compounds inhibit the catalytic activity of aromatase and as a consequence lead to a number of undesirable side-effects, including arthralgia and possible cognitive defects due to inhibition of aromatase in the bone and brain, respectively. Novel therapies, such as those employed to treat obesity-associated disease, including anti-diabetics, may prove successful at inhibiting aromatase specifically within the breast. This SpringerBrief will explore all of these issues in depth and the authors are in a unique position to write about this topic, having extensive experience in the field of aromatase research.?

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