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system will be used to measure, with high sensitivity, catecholamines, steroid hormones, and their synthesizing enzymes. In addition, we will use sophisticated imaging techniques in the in vitro and in vivo systems to demonstrate functional relationships between adrenergic and sex steroid synthesis in normal and cancerous breast tissue. The project has two aspects: 1) We will determine the relative contribution of catecholamines to estrogen and progesterone synthesis by normal human breast tissue in the in vitro culture system. 2) Using the in vitro model of human breast tissue, we will test the effects of catecholamines on estrogen and progesterone secretion by normal and cancerous human breast tissue. We will determine how adrenergic stimulation of estrogen and progesterone synthesis is affected by the addition of E2 to the culture medium, by pulsing cultures with gonadotropins, by addition of antiprogestins, by interruption of adrenergic action, and by inhibition of aromatase. The regulation of E2 and P4 synthesis by catecholamines in normal tissue will be compared to that in cancerous tissue and the pattern of regulation will be compared to that in normal women. The information we obtain from these studies will lead to a better understanding of the mechanisms of breast cancer, and may identify a therapeutic pathway to treat and prevent this disease.

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