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Corticosterone content in rat mammary cells is reduced by the aldosterone antagonist spironolactone but not by the 5 alpha-reductase inhibitor finasteride. The adrenal steroids aldosterone and corticosterone have opposing effects on mammary gland development. Aldosterone is known to enhance lobuloalveolar differentiation while corticosterone tends to block this development. Finasteride, an inhibitor of 5 alpha-reductase that reduces the plasma level of dihydrotestosterone (DHT) and thereby elevates the plasma level of corticosterone, was used to study the role of corticosterone in regulating lobuloalveolar differentiation. Female rats were ovariectomized, treated

with estradiol and infused intravenously with 500 microg estradiol-17 beta (E2) +/-day progesterone (P4) for 10 days. On day 5, a group of rats received either 0.5 mg finasteride/day or vehicle in addition to E2 +/-day P4 for the next 9 days. Mammary glands were then removed for morphological and biochemical studies. Incorporation of [3H]thymidine into mammary DNA was used to determine the extent of cell proliferation. Dihydrotestosterone (DHT) content was quantitated in mammary tissue to detect any alterations in the availability of androgen to the tissue. The steroid content of mammary cells was determined by measuring their corticosterone content. Finasteride had no effect on the extent of proliferation of mammary epithelial cells and on the content of corticosterone or DHT in the mammary

tissue. The apparent lack of an effect on mammary steroid levels is consistent with our previous study showing that DHT alone does not stimulate cell proliferation. When finasteride was administered, lobuloalveolar differentiation was reduced. However, no effect of finasteride on mammary steroid content was observed. We conclude that finasteride, by reducing the availability of DHT to the mammary gland, can stimulate lobuloalveolar differentiation. This response appears to be independent of alterations in the levels of corticosterone in the gland.

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