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## Inhibitory Effect of 15-Oxygenated Sterols on Cholesterol Synthesis from 24, 25-Dihydrolanosterol\*

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Several 15-oxygenated sterols were examined as to their inhibitory activity toward cholesterol synthesis from [24,25- $^3$ H]-24,25-dihydrolanosterol in the 10,000×g supernatant fraction of a rat liver homogenate (Table 1). At 40  $\mu$ M, three 15 $\alpha$ -hydroxylated compounds, 14 $\alpha$ -ethylcholest-7-ene-3 $\beta$ ,15 $\alpha$ -diol, 14 $\alpha$ -methylcholest-7-ene-3 $\beta$ ,15 $\alpha$ -diol, and lanost-7-ene-3 $\beta$ ,15 $\alpha$ -diol, were found to be extremely potent inhibitors (more than 90% inhibition) of dihydrolanosterol metabolism. The inhibitory effect of the C-15 substituents appears to be in the order of : 15 $\alpha$ -hydroxyl>15-ketone>15 $\beta$ -hydroxyl.

Table I. Cholesterol biosynthesis during incubation of the S-10 fraction of a rat liver homogenate with [24,25-3H]-24,25-dihydrolanosterol in the presence of oxygenated sterols.

	Distribution of radioactivity in		T 1 *1 *4 *
Compound	Dihydrolanosterol fr. (%)	Cholesterol (%) fr. (%)	Inhibitior (%)
None (Control)	32,0	31,2	
Cholest-8(14)-ene-3 $\beta$ ,15 $\beta$ -diol	32, 1	29.7	5
Cholest-(14)-en-3 $\beta$ -ol-15-one	49.3	18.1	42
$14\alpha$ -Methylcholest-7-en-3 $\beta$ -ol-15-one	51,8	16, 2	48
$14\alpha$ -Methylcholest-7-ene- $3\beta$ , $15\alpha$ -diol	89.9	1.3	96
$14\alpha$ -Ethylcholest-7-en-3 $\beta$ -ol-15-one	66, 3	8.8	72
$14\alpha$ -Ethylcholest-7-ene- $3\beta$ , $15\alpha$ -diol	88,5	2,3	93
$14\alpha$ -Ethylcholest-7-ene- $3\beta$ , $15\beta$ -diol	63, 1	10.0	68
Lanost-7-en-3 $\beta$ -ol-15-one	85.6	4.2	87
Lanost-7-ene-3 $\beta$ ,15 $\alpha$ -diol	88.5	1,2	96
Lanost-7-ene-3 $\beta$ ,15 $\beta$ -diol	55.9	15.3	51
15ξ-Fluorolanost-7-en-3β-ol	59.4	12,3	61
Lanost-7-en-3β-ol	39.6	25,6	18

<sup>\*</sup> 本報告は J. Biochem., 99, 597-600 (1986) に発表.

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