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Author	荻原, 典子(Ogihara, Noriko) 森崎, 益雄(Morisaki, Masuo)
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Facile Synthesis of Zymosterol and Related Compounds*

Noriko Ogihara and Masuo Morisaki

荻原典子, 森崎益雄

Facile preparations of cholesta-8,24-dien-3 β -ol, zymosterol (1) and the related 7,24-diene (2), 5,7,24-triene (3) and 8,14,24-triene (4), all of which are potential intermediates of cholesterol or ergosterol biosynthesis, are described.

5-Cholene-3 β ,24-diol 3-tetrahydropyranyl ether is oxidized to the 24-aldehyde, which is then converted into the 3β -acetoxy-24-acetal (5) mp, $124-126^{\circ}$ C through acetalization and acetylation. Successive treatment of 5 with N-bromosuccinimide, tetra-n-butylammonium bromide and tetra-n-butylammonium fluoride gives chola-5,7-diene-3 β -tetrahydropyranyloxy-24-acetal (6, 43% yield), the common synthetic progenitor of all the targeted sterols. Isomerization of 6 with p-toluenesulfonic acid gives the 8,14-diene (7) in 55% yield, after basic hydrolysis. Catalytic hydrogenation of the 5,7-diene (6) and 8,14-dien-3 β -ol (7) gives the 7-ene (8), or a mixture (ca. 1:1) of the 8(9)-ene (9) and the 8(14)-ene (10), respectively.

Deacetalization of the corresponding acetates of 9 plus 10 followed by Wittig reaction with isopropylidene triphenylphosphorane yields, after saponification, a mixture (ca. 1:1) of zymosterol (1), and cholesta-8(14), 24-dien-3 β -ol in 40% yield. Recrystallization of this material from methanol gave zymosterol, mp 109-111°C. In the same manner, deacetalization of 6, 7, and 8 followed by Wittig reaction gives the 5,7,24-triene (3), 8,14,24-triene (4) and 7,24-diene (2), respectively.

HPLC and GC behaviors of these sterols, which would be useful for their identification in biological systems, are also described.

^{*} 本報告は Chem. Pharm. Bull., 36, 2724 (1988) に発表.