## A VOLATILE METABOLITE OF ACTINOMYCETES, 2-METHYLISOBORNEOL

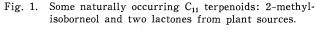
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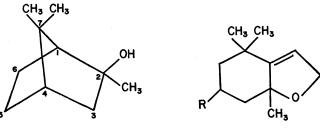
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(Received for publication August 25, 1969)

During the past five years we have isolated geosmin, trans-1, 10-dimethyl-trans-9-decalol from 20 different actinomycetes<sup>1)</sup>. However, from 4 of these actinomycetes the major volatile component was a crystalline solid with a strong camphor-menthol-like odor. They were Streptomyces lavendulae IMRU 3440-1 Y, S. sp. 100-1 (received from JACOB EREN, isolated from a reservoir in Israel), S. sp. AY-219 (received from BENJAMIN BECKER, isolated from soil, Clinton, N.Y.), and Actinomadura sp. I-15 (received from M. P. LECHEVALIER, isolated from greenhouse soil). The nuclear magnetic resonance and mass spectra of this solid suggested that it was 2-methylisoborneol (current Chem. Abstr. name: 1,2,7,7-tetramethyl-2-norbornanol). The identity was proved by comparison with an authentic sample prepared from camphor and methylmagnesium iodide<sup>2)</sup>.

At this point, others reported that 2methylisoborneol is the major odorous component of *S. antibioticus* 5324, *S. praecox* ATCC 3374 and *S. griseus* ATCC 10137<sup>8</sup>). Our work confirms this identification and gives additional useful data.



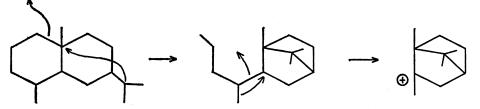


R=H, OH

The melting point of 2-methylisoborneol is 170°C (sealed tube), lit. mp. 168°C<sup>2)</sup>. The mass spectrum shows bands in decreasing order of intensity at 107, 95, 41, 93, 43, 79, 55, 91, 94, 135, 108, 150, 77 mass units. The differences from those reported by MEDSKER, et al. are qualitative, to be expected when comparing mass spectra from different instruments run at different temperatures. More important, the structure of 2-methylisoborneol can be deduced from published generalizations resulting from a comprehensive mass spectral study of norbornane derivatives<sup>4)</sup>. For sample: (A) a strong band at 41 indicates a gem-dimethyl group, (B) weak bands at 57 and 71 mean no gem-dimethyl group in position 3 (as in fenchyl alcohol), (C) a strong band at 95 indicates 2 or 3 (depending upon mono or di-substitution at position 7) methyl groups at positions 1 and 4 thru 7, (D) a strong band at 58 (compared to 44, 72 and 86) means a methyl substituent at position 2 not 3. The four singlet methyl absorption bands in the nmr spectrum show significant differences in line width which are due to long range spin-spin couplings characteristic of this ring system<sup>5)</sup>.

Borneol has been isolated from some fermentations to which  $\alpha$ -pinene or camphene had been added<sup>6)</sup>. However, 2methylisoborneol was previously unknown in nature. Eleven-carbon terpenoids are rare; the two lactones in Fig. 1 have each been isolated from several plants<sup>7)</sup> and trans-2,2,6-trimethylcyclohexylacetic acid was found in petroleum<sup>8)</sup>. Most natural terpenoids with an unusual number of carbon atoms seem to be derived biosyn-

thetically from a larger, normal (*i. e.*  $C_{10}$ ,  $C_{15}$ ,  $C_{20}$ ) terpenoid by loss of carbon. Examples are cryptone<sup>30</sup>, geosmin, cogejerine<sup>10)</sup>, and gibberellic acid. Thus, 2methylisoborneol might be biosynthesized as shown in Fig. 2 although, superficially it appears to be a typical monoterpene unit (borneol) plus a methyl group. Fig. 2. Possible biosynthesis of 2-methylisoborneol from a typical sesquiterpene precursor.



## Acknowledgement

We thank ED EMERY and the Colgate-Palmolive Research Center for the mass spectrum, E. M. FEKETE for technical assistance. This investigation was supported by Public Health Service Grant AI 06708 from the National Institute of Allergy and Infectious Diseases.

## Literature cited

- GERBER, N. N.: Geosmin, from microorganisms, is trans-1,10-dimethyl-trans-9-decalol. Tetrahedron Letters 1968-25: 2971~ 2974, 1968.
- TOIVONEN, H.: Über die Reaktionen des 2-Methylisoborneols und des 2-Methylfenchols in konzentrierter Salpetersäure. Tetrahedron Letters 1968-26: 3041~3044, 1968.
- 3) MEDSKER, L. L.; D. JENKINS & J. F. THOMAS: Odorous compounds in natural waters. 2-Exo-hydroxy-2-methylbornane, the major odorous compound produced by several actinomycetes. Environmental Science & Technology 3: 476~477, 1969.
- DIMMEL, D. R. & J. WOLINSKY: An electronimpact study of norbornane derivatives. J. Org. Chem. 32: 2735~2739, 1967.

- 5) BAKER, K. M. & B. R. DAVIS: Long range coupling and solvent shifts in the nmr spectra of bornane derivatives. Tetrahedron. 24: 1663~1672, 1968.
- SHUKLA, O. P.; M. N. MOHOLAY & P. K. BHATTACHARYYA: Microbiological transformations of terpenes. X. Fermentation of α- and β-pinenes by a soil pseudomonad (PL-strain). Indian J. Biochem. 5:79~91, 1968.
- BAILEY, W. C., Jr.; A. K. BOSE, R. M. IKEDA, R. H. NEWMAN, H. V. SECOR & C. VARSEL: The isolation from tobacco of 2-hydroxy-2, 6,6-tri-methylcyclohexylidene acetic acid γlactone and its synthesis. J. Org. Chem. 33: 2819~2822, 1968.
- CASON, J. & K. L. LIAUW: Characterization and synthesis of a monocyclic eleven-carbon acid isolated from a California petroleum. J. Org. Chem. 30: 1763~1769, 1965.
- COFFEY, S. (Editor): RODD's chemistry of carbon compounds. 2 nd Edition, Vol. II, part B, p. 200. Elsevier Publishing Co., New York, 1968.
- 10) GOUGH, J.; V. POWELL & M. D. SUTHERLAND: Constitution and biogenesis of two new sesquiterpenes. Tetrahedron Letters 1961 -21: 763~767, 1961.