

These data suggested that the compound might be the dihydro-derivative of 9,10-dihydroxy- Δ^6 _{a(10a)}-tetrahydrocannabinol. Assuming that the stereochemistry at C 6a and C 10a is the same as in the naturally occurring Δ^9 -THC, four stereoisomers should be considered.

Synthesis of these isomers showed that (–) (6aR, 9S, 10S, 10aR)-9,10-dihydroxy-hexahydrocannabinol was identical (¹H-NMR, IR, MS, UV, R_p [α]_D) with the isolated compound. The synthesis of **I** was accomplished by epoxidation of (–) (6aR, 10aR)- Δ^9 -THC-acetate followed by alkaline hydrolysis and isolation by repeated chromatog-

raphy of the reaction mixture. This is the first report of the isolation of (–) (6aR, 9S, 10S, 10aR)-9,10-dihydroxy-hexahydrocannabinol from *Cannabis* which we named cannabiripsol.

Due to the stereochemistry of cannabiripsol and other cannabinoids such as Δ^9 -THC the 'activity' of cannabiripsol should more nearly mimic that of Δ^9 -THC type cannabinoids than that of other dihydroxylated cannabinoids. The pharmacological profile of cannabiripsol is currently being investigated.

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- 9 The synthesis and spectral characteristics of all isomers as well as full details on the isolation of (–) (6aR, 9S, 10S, 10aR)-9,10-dihydroxyhexahydrocannabinol will be described elsewhere.

(–)-Maalian-5-ol, a new enantiomeric sesquiterpenoid from the liverwort *Plagiochila ovalifolia*

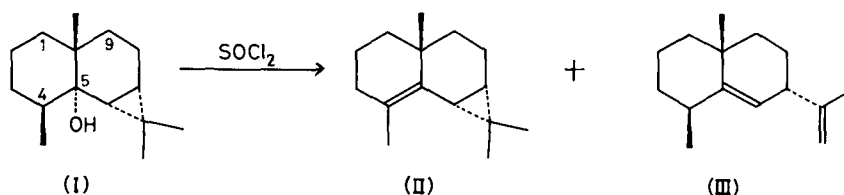
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Summary. A new enantiomeric sesquiterpene alcohol named (–)-maalian-5-ol was isolated from the liverwort, and the structure and absolute configuration was determined to be the stereostructure **I** by chemical and spectral evidence.

In the course of our investigation on terpenoids from the liverworts (Hepaticae), which form a unique group in the plant kingdom, several enantiomeric sesquiterpenoids² being antipodes to the normal ones from vascular plants were isolated along with several novel carbon skeletal sesquiterpenoids³. The present communication deals with the isolation and the structure determination of an additional new enantiomeric sesquiterpene alcohol, named (–)-maalian-5-ol, from the leafy liverwort, *Plagiochila ovalifolia* Mitt. belonging to the Plagiochilaceae in the Jungermanniaceae. The liverwort (860 g), collected at Kuju in Kyushu and dried in the shade for a few days, was digested with methanol. The methanol solution, after being concentrated in vacuo, was then extracted with ether and the solvent was distilled out at reduced pressure to afford a viscous matter (10.2 g). Part of the extract (6.8 g) was chromatographed over silica gel with a mixed solvent of chloroform and ether (v/v, 5:1) to isolate the new sesquiterpene alcohol (150 mg) as a major constituent. Spectroscopic evidence showed that the compound, C₁₅H₂₄O (M⁺ 222.1990), [α]_D²⁰ + 106° (c 2.5, CHCl₃), was a saturated tricyclic sesquiterpene alcohol containing a cyclopropane ring [δ_{CDCl₃} 0.4–0.8 (2H, com-

plex)] together with a tertiary hydroxyl group [ν_{CCl₄} 3625 and 3500 cm^{–1}; no signal between δ 3.0 and 4.5], a secondary methyl [δ 0.93 (3H, d, J=7.5)] and 3 tertiary methyls [δ 1.00, 1.05 and 1.10 (each 3H, s)]. This structure was supported by the off-resonance ¹³C-NMR spectrum which showed 3 singlets (δ 17.6, 36.4 and 75.0), 3 doublets (δ 20.4, 32.9 and 34.1), 5 triplets (δ 16.9, 21.6, 30.0, 31.0 and 37.0) and 4 quartets (δ 16.5, 16.5, 23.4 and 31.0). The alcohol (120 mg) was dehydrated with SOCl₂ in pyridine to yield 2 kinds of hydrocarbons, which were, respectively, isolated as a less polar (40 mg) and a more polar (40 mg) product by means of preparative TLC over silica gel using hexane as a solvent. The spectral properties of the former (**II**), C₁₅H₂₄ (M⁺ 204); ν_{CCl₄} 1230, 1140, 1070, 973 and 953 cm^{–1}; δ_{CDCl₃} 0.83, 0.99 and 1.09 (each 3H, s), 1.57 (3H, br.s), coincided with those of (–)-β-maaliene reported by Büchi et al.⁴, and the latter (**III**), C₁₅H₂₄ (M⁺ 204); ν_{CCl₄} 3060, 1650, 1210, 985, 955, 900 and 885 cm^{–1}; δ_{CDCl₃} 1.15 (3H, d, J=8.0), 1.14 (3H, s), 4.65 and 4.78 (each 1H, br.s), 5.20 (1H, d, J=4.0), did with those of (+)-selina-5,11-diene⁴. However, the optical rotations of both the sesquiterpene hydrocarbons (**II**, [α]_D²⁰ + 129° and **III**, [α]_D²⁰ – 145°) derived from



(-)-maalian-5-ol (**I**) gave opposite sign compared with the values ($[\alpha]_D -138^\circ$ and $[\alpha]_D +149^\circ$) of the known compounds⁴.

Accordingly, it was certain that the alcohol consists of the structure and absolute configuration, excluding the tertiary hydroxyl group on C-5, as shown in the stereostructure **I**. Next, the configuration of the hydroxyl group was elucidated by examination of the solvent effect on ¹H-NMR spectra to be trans to both the methyls on C-4 and C-10: when the ¹H-NMR spectra of the alcohol (**I**) were measured in CDCl₃ and C₅D₅N solutions, the C-4 secondary methyl (Δ 0.04) and the C-10 tertiary methyl (Δ 0.02)

exhibited no solvent shift due to vicinal deshielding of the C-5 hydroxyl group⁵.

On the basis of the above chemical and spectroscopic evidence, the structure and absolute configuration of the sesquiterpene alcohol, (-)-maalian-5-ol, was determined to be shown by the stereostructure **I**. This may be biosynthesized from trans-farnesyl pyrophosphate via (-)-bicyclogermacrene which has been isolated as a common component in the liverworts⁶. We were interested in such occurrence of the enantiomeric sesquiterpenoids in the liverworts with respect to a chemotaxonomy of the plants and a biogenesis of the compounds.

- 1 The authors express their gratitude to Dr H. Inoue, National Science Museum, for identification of the liverwort, and Prof. T. Tokoroyama, Osaka City University, for sending of the data of (-)- β -maaliene and (+)-selina-5,11-diene. Thanks are also due to the Ministry of Education for financial support of this work.
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Biochemical compartmentation of fish tissues. II. Nonspecific phosphomonoesterases in brain

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Summary. The specific activities of acid and alkaline phosphatases in different regions of the brain of 9 nutritionally important fishes were worked out. The region consisting of pituitary, hypothalamus and thalamus showed the highest acid and alkaline phosphatase activities. Least activities of both the enzymes were found in the cerebellum and medulla oblongata. The piscivorous fishes contain the highest acid and alkaline phosphatase activity followed by cat fishes and major carps. The distributional pattern of these 2 enzymes in 4 regions of the brain of 9 fishes is the same.

Recently, studies in the biochemical aspect of animal tissues has been encouraged for a better insight into biochemical organisation²⁻⁴. Much attention has been paid in the past to the mammalian brain with regard to key enzymes like glutamate de-carboxylase, choline acetyltransferase, glucose-6-phosphatase, fructose 1, 6-diphosphatase, important neurotransmitters like catecholamines, energetic substances like adenine-nucleotides, glycogen, glucose, lactate and pyruvate⁵⁻¹⁰. However, the compartmentation of the brain in fishes has to be investigated. It is known that various parts of the brain differ in their biochemical make-up following distinct phylogenetic ages. Although acid and alkaline phosphatases are widely distributed in animal tissues, they are mainly concerned with the reabsorption of glucose molecules in renal tissue, to conduct an active glycolytic mechanism in liver, dephosphorylation of glucoesters in sex organs, active involvement in protein synthesis, association with the secretory activity and absorption of nutrients in digestive system, phosphate cycles and growth¹¹⁻¹³. However, the functional significance of these phosphatases in various regions of the brain is still to be completely known. If we want to understand fully the complex functions of the brain as a central, controlling, regulating and co-ordinating organ for sensory and memo-

ry functions, it is essential to know the levels of the non-specific phosphomonoesterases in different regions of brain, because of the close relation between brain metabolism and neuronal activity¹⁴.

The functional significance of alkaline phosphatase (orthophosphoric monoester phosphohydrolase EC 3.1.3.1) in brain is supposedly involved in various secretory and transport processes and in blood-brain barrier mechanisms¹⁵. The acid phosphatase (orthophosphoric monoester phosphohydrolase EC 3.1.3.2) is implied again in the secretory processes and in the intracellular transport mechanisms. Such non-specific phosphomonoesterases are also involved in transphosphorylation³. The main aim of the present investigation is to study the differential distribution and functional role of these phosphatases in 4 regions of the brain i.e. cerebrum, cerebellum, the region consisting of pituitary, hypothalamus and thalamus and medulla oblongata of 9 nutritionally and economically important fishes i.e. *Labeo rohita* (Ham), *Cirrhina mrigala* (Ham), *Catla catla* (Ham) (herbivorous and soft fishes), *Channa punctatus* (Bloch), *Channa marulius* (Ham), *Channa striatus* (Bloch) (hard and piscivorous) and *Clarias batrachus* (Linn), *Heteropneustes foliolis* Bloch and *Mystus seenghala* (sykes) (hard and carnivorous). This is a first