

OXOANLOBINE, A NEW OXOAPORPHINE ALKALOID FROM GUATTERIA MELOSMA

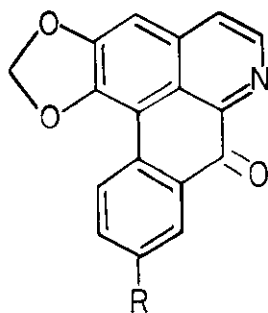
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Abstract - Oxoanolobine (1), a new alkaloid from an alcoholic extract of Guatteria melosma Diels (Anonaceae), was characterized as 1,2-methylenedioxy-9-hydroxyoxoaporphine (10-hydroxy-8H-benzo[g]-1,3-benzodioxolo[6,5,4-de]quinolin-8-one) by physicochemical data and conversion to lanuginosine (2).

An alcoholic extract (654 g) of Guatteria melosma¹ (27.2 kg) was submitted to an acid-base partition procedure. Subsequent silicic acid chromatography of the basic fraction afforded oxoanolobine (1) (50 mg) as an orange amorphous solid from methanol; mp 270-275° (dec); $[\alpha]_D^{28} 0^\circ$ (c 1.0, MeOH);
 uv $\lambda_{\max}^{\text{MeOH}}$ nm 217 (log ϵ 4.24) 249 (4.43), 274 (4.35), 324(sh) (3.84), 370 (3.65) and 442 (3.76),
 $\lambda_{\max}^{\text{MeOH} + 0.1N \text{ HCl}}$ nm 220 (log ϵ 4.29) 260 (4.41), 287 (4.31), 345 (3.72), 395 (3.73) and 510 (3.51);
 $\lambda_{\max}^{\text{MeOH} + 0.1N \text{ NaOH}}$ nm 222 (log ϵ 4.29), 253 (4.38), 291 (4.42), 334(sh) (4.01), 372 (3.48) and 506 (3.42); ir ν_{\max}^{KBr} cm^{-1} 3420 (br), 1660 (C=O). The nmr spectrum (60 MHz, TFA, TMS, δ in ppm) indicated the presence of one methylenedioxy group at 6.60 (2H,s), a C-3 aromatic proton at 7.47 (1H,s) an aromatic AB system for C-4 at 8.37 (1H,d,J=6Hz) and C-5 at 8.65 (1H,d,J=6Hz) and an aromatic AMX system for C-8, C-10 and C-11 at 7.98 (1H,d,J=2.5Hz), 7.59 (1H,dd,J=8,2.5Hz) and 8.70 (1H,d,J=8Hz), respectively. The ms showed a M⁺ at m/e 291 (100%) for C₁₇H₉NO₄, 263(8), 233(15) and 178(10) with metastable ions at m/e 237.3 for the transition 291→263 (m*_{calc} 237.7) and 206.4 for the transition 263→233 (m*_{calc} 206.4). These spectral data indicated that 1 was 1,2-methylenedioxy-9-hydroxyoxoaporphine.

Treatment of oxoanolobine (1) with ethereal diazomethane gave an O-methyl derivative (lanuginosine) (2) as yellow needles from methanol; mp 314°; $[\alpha]_D^{28} 0^\circ$ (c 1.0, MeOH); ms M⁺ m/e 305(100%), 304(56), 290(5), 276(20), 275(40) and 234(16). O-Methylxoanolobine was identical (ir, uv, ms, mp) with authentic lanuginosine², thus confirming that 1 was 1,2-methylenedioxy-9-hydroxyoxoaporphine (10-hydroxy-8H-benzo[g]-1,3-benzodioxolo[6,5,4-de]quinolin-8-one)³.



- 1 R = OH
 2 R = OCH₃
 3 R = H

Oxoaporphine alkaloids have been found in several plant families in addition to the Anonaceae. These include the Araceae, Hernandiaceae, Lauraceae, Magnoliaceae, Menispermaceae, Monimiaceae, Papaveraceae, Ranunculaceae and Eupomatiaceae.^{4,5} Although liriodenine (3) has reported broad spectrum antimicrobial activity^{6,7} and activity against 9-KB tissue culture cells⁸, little is known about the pharmacological activity of other members of this group of alkaloids.

Acknowledgements and References

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4. M. Shamma, "The Isoquinoline Alkaloids", Academic Press, New York, 1972, p. 245.
5. M. Shamma and J.L. Moniot, "Isoquinoline Alkaloids Research", Plenum Press, New York, 1978, p. 173.
6. C.R. Chen, J.L. Beal, R.W. Doskotch, L.A. Mitscher and G.H. Svoboda, *Lloydia*, 1974, 37, 493.
7. C.D. Hufford, M.J. Funderburk, J.M. Morgan and L.W. Robertson, *J. Pharm. Sci.*, 1975, 64, 789.
8. P.E. Sonnet and M. Jacobson, *J. Pharm. Sci.*, 1971, 60, 1254.

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