

BIBLIOGRAPHIE

1. DuVigneaud, P. (1952) *Bull. Soc. Roy. Bot. Belg.* **85**, 9.
2. Denoël, A. et Jaminet, F. (1953) *Contribution à l'étude chimique des Strychnos du Congo Belge*, Ministère des Colonies, eds. Bruxelles.
3. Turkovic, I. (1968) *J. Pharm. Belg.* **23**, 283.
4. Bouquet, A. (1972) *Plantes Médicinales du Congo-Brazzaville* (O.R.S.T.O.M. ed.). Paris.
5. Klyne, W., Swan, R. J., Bycroft, B. W. et Schmid, H. (1966) *Helv. Chim. Acta.* **49**, 833.
6. Wenkert, E. et Sklar, R. (1966) *J. Org. Chem.* **31**, 2689.
7. Hymon, J. R. et Schmid, H. (1966) *Helv. Chim. Acta* **49**, 2067.
8. Bosly, J. (1951) *J. Pharm. Belg.* [n.s.] **6**, 150 et 243.

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ALKALOIDS OF *CISSAMPELOS PAREIRA**

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Plant. *Cissampelos pareira* L. (Menispermaceae), a woody climber of the forests of Ghana. The plant was collected and identified by Mr. K. Obeng-Darko, of the Faculty of Agriculture, University of Science and Technology, Kumasi. Voucher specimens are on deposit at the Faculty of Pharmacy of the University. *Uses.* Medicinal [1]. *Previous work.* On the roots [2-14], whole plant [15,16] and the leaves [17] for the alkaloids hayatine [(±)-bebeerine] [(±)-curine] [2,6,8-11,17], hayatinine [(-)-4"-O-methylbebeerine] [(-)-4"-O-methylcurine] [2,5,7-9,12,17], (-)-bebeerine [4-6,8-10,17], (+)-isochondodendrine [6,8-10], cyclanoline chloride (cissamine chloride) [9,14], cissampareine [15,16], (+)-4"-O-methylcurine [(+)-4"-O-methylbebeerine] [18], hayatidine [(±)-4"-O-methylbebeerine] [(±)-4"-O-methylcurine] [13,17], cycleanine [17] and for (+)-quercitol [2-5,17], sterols [2,4,5] and essential oil [4,5].

Plant part examined. The powdered roots (2.25 kg) were percolated with dil HOAc (10%). The acidic extract (39 l.) was basified with NH₄OH and extracted with CHCl₃ (49 l.) in a batchwise fashion. The combined CHCl₃ extracts were dried (Na₂SO₄), filtered and the filtrate evaporated *in vacuo* to afford an alkaloidal residue (39 g). Adsorption of this residue on Al₂O₃ (36 g) followed by chromatography over Al₂O₃ (88 g) in Et₂O and then Et₂O-MeOH afforded the following alkaloids (in order of elution):

Dehydrodicentrine. 26 mg; mp 216° (Et₂O); $[\alpha]_D^{21}$ 0° (CHCl₃; c 0.35); $\lambda_{\max}^{\text{MeOH}}$ nm (log ϵ): 217 (4.40), 266 (4.87), 303 (4.02) and 339 (4.23); $\nu_{\max}^{\text{KBr}}$ cm⁻¹: 1595, 1515, 1470, 1397, 1308, 1209, 1114 and 857; $\delta_{60\text{ MHz}}^{\text{CDCl}_3}$: 3.08 (s, 3H, NMe), 3.99 (s, 6H, OMe), 6.12 (s, 2H, CH₂O₂), 6.47 (s, 1H, ArH), 6.82 (s, 1H, ArH), 6.99 (s, 1H, ArH) and 8.34 (s, 1H, ArH); MS M⁺ m/e 337 (100%) for C₂₀H₁₉NO₄, 322 (88), 293 (8), 279 (7), 251 (4) and 168.5 (18) identical by direct comparison (UV, IR, NMR, mp, mmp) with an authentic sample.

Dicentrine. 412 mg; mp 167-168.5° (Et₂O); $[\alpha]_D^{21}$ +57.77° (CHCl₃; c 1.35); $\lambda_{\max}^{\text{MeOH}}$ nm (log ϵ): 220 (4.21), 281 (3.88), 305 (3.92) and 315 (sh) (3.88); $\nu_{\max}^{\text{KBr}}$ cm⁻¹: 1603, 1520, 1461, 1455, 1390, 1341,

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1267, 1245, 1214, 1097, 928, 865, 770 and 608; $\delta_{60\text{MHz}}^{\text{CDCl}_3}$: 2.53 (s, 3H, NMe), 3.90 (s, 6H, OMe), 6.84 (d) and 6.98 (d) (Jgem 2Hz) (2H, CH₂O₂), 6.42 (s, 1H, ArH), 6.69 (s, 1H, ArH), and 7.56 (s, 1H, ArH); MS M⁺ m/e 339 (69%) for C₂₀H₂₁NO₄, 338 (100), 337 (7), 324 (7), 322 (7), 309 (9), 296 (17), 265 (12) and 238 (4) identical by direct comparison (UV, IR, NMR, MS, mp, mmp) with an authentic sample.

Cycleanine. 75 mg; mp 280° (Et₂O); $[\alpha]_D^{22}$ -15.94° (CHCl₃; c 0.69); $\lambda_{\text{max}}^{\text{MeOH}}$ nm (log ϵ): 232 (sh) (4.87), 276 (3.89) and 285 (sh) (3.83); $\nu_{\text{max}}^{\text{KBr}}$ cm⁻¹: 1608, 1505, 1490, 1452, 1415, 1375, 1340, 1293, 1220, 1170, 1145, 1115, 1068, 1017, 843 and 805; $\delta_{60\text{MHz}}^{\text{CDCl}_3}$: 2.53 (s, 6H, NMe), 3.38 (s, 6H, OMe), 3.78 (s, 6H, OMe), 6.13-7.05 (m, 10H, ArH); MS M⁺ m/e 622 (29%) for C₃₈H₄₂N₂O₆, 621 (8), 313 (26), 312 (100), 311 (28), 204 (29), 190 (17), 174 (17), 159 (14), 146 (7) and 145 (8) identical by direct comparison (UV, IR, NMR, MS, mp, mmp) with an authentic sample.

Insularine. 45 mg; mp 157° (Et₂O); $[\alpha]_D^{22}$ +11.36° (EtOH; c 0.44); $\lambda_{\text{max}}^{\text{EtOH}}$ nm (log ϵ): 209 (4.68), 229 (sh) (4.56) and 276 (3.64); $\nu_{\text{max}}^{\text{KBr}}$ cm⁻¹: 1605, 1580, 1500, 1450, 1420, 1365, 1260, 1215, 1114, 1070, 1010, 834 and 810; MS M⁺ m/e 620 (90%) for C₃₈H₄₀N₂O₆, 619 (38), 605 (6), 313 (20), 312 (100), 311 (28), 310 (74), 309 (26), 204 (14), 190 (12), 174 (9), 159 (8), 146 (8) and 145 (9) identical by direct comparison (UV, IR, MS, mp) with an authentic sample.

Isochondodendrine. 78 mg; mp 295° (Et₂O-MeOH); $[\alpha]_D^{22}$ +40.32° (pyr; c 0.62); $\lambda_{\text{max}}^{\text{EtOH}}$ nm (log ϵ): 211 (4.72), 231 (sh) (4.58), 278 (3.73) and 285 (3.72); $\nu_{\text{max}}^{\text{KBr}}$ cm⁻¹: 1610, 1585, 1508, 1450, 1295, 1215, 1170, 1115, 1065, 1042, 1015, 975, 940, 928, 855, 845, 815, 805 and 759; MS M⁺ m/e 594 (60%) for C₃₆H₃₈N₂O₆, 593 (14), 487 (4), 312 (4), 299 (14), 298 (100), 297 (20), 191 (9), 190 (7) and 162 (6) identical by direct comparison (UV, IR, MS, mp, mmp) with an authentic sample.

Although isochondodendrine has been isolated several times from *C. pareira* [6,8-10], cycleanine has only recently been isolated from *C. pareira* [17] and from *C. insularis* [20]. Furthermore, this is the first reported isolation of dehydrodicentrine and dicentrine in the genus *Cissampelos* and the first reported isolation of insularine in *C. pareira*, the latter base first isolated from *C. insularis* [21].

Biological significance. Alkaloids of the plant have pronounced pharmacological effects [10,22].

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REFERENCES

1. Dwuma-Badu, D. (1974) Personal communication.
2. Bhattacharji, S., Sharma, V. N. and Dhar, M. L. (1952) *J. Sci. Ind. Res.* **11B**, 81.
3. Council for Scientific and Industrial Research (Indian) (1955) 50340 *Chem. Abstr.* **50**, 538.
4. Bhattacharji, S., Sharma, V. N. and Dhar, M. L. (1955) *Bull. Natl. Inst. Sci. India* No. 4, 39; *Chem. Abstr.* **50**, 2626.
5. Bhattacharji, S., Sharma, V. N. and Dhar, M. L. (1956) *J. Sci. Ind. Res.* **15B**, 363.
6. Kupchan, S. M., Yokoyama, N. and Beal, J. L. (1960) *J. Am. Pharm. Assoc. Sci. Ed.* **49**, 727.
7. Bhattacharji, S., Roy, A. C. and Dhar, M. L. (1962) *J. Sci. Ind. Res.* **21B**, 428.
8. Srivastava, R. M. and Khare, M. P. (1963) *Current Sci. (India)* **32**, 114.
9. Srivastava, R. M. and Khare, M. P. (1964) *Chem. Ber.* **97**, 2732.
10. Boissier, J. R., Combes, G., Pernet, R. and Dumont, C. (1965) *Lloydia* **28**, 191.
11. Bhatnagar, A. K., Bhattacharji, S., Roy, A. C., Popli, S. P. and Dhar, M. L. (1967) *J. Org. Chem.* **32**, 819.
12. Bhatnagar, A. K. and Popli, S. P. (1967) *Ind. J. Chem.* **5**, 102.
13. Bhatnagar, A. K. and Popli, S. P. (1967) *Experientia* **23**, 242.
14. Anwer, F., Popli, S. P., Srivastava, R. M. and Khare, M. P. (1968) *Experientia* **24**, 999.
15. Kupchan, S. M., Patel, A. C. and Fujita, E. (1965) *J. Pharm. Sci.* **54**, 580.
16. Kupchan, S. M., Kubota, S., Fujita, E., Kobayashi, S., Block, J. H. and Telang, S. A. (1966) *J. Am. Chem. Soc.* **88**, 4212.
17. Chowdhury, A. R. (1972) *Sci. Cult.* **38**, 358.
18. Haynes, L. J., Herbert, E. J. and Plimmer, J. R. (1966) *J. Chem. Soc. (C)* **1966**, 615.
19. Cava, M. P., Watanabe, Y., Bessho, K., Mitchell, M. J., Darocha, A. I., Hwang, B., Douglas, B. and Weisbach, J. A. (1968) *Tetrahedron Letters* **1968**, 2437.
20. Kondo, H., Tomita, M. and Uyeo, S. (1937) *Chem. Ber.* **70B**, 1890; *Chem. Abstr.* **31**, 8539.
21. Kondo, H. and Yano, K. (1927) *J. Pharm. Soc. Japan* No. 548, 815; *Chem. Abstr.* **22**, 786.
22. Patnaik, G. K., Pradhan, S. N. and Vohra, M. M. (1973) *Indian J. Exp. Biol.* **11**, 89; *Chem. Abstr.* **80**, 103855.