Alkaloids from the Roots of Saccopetalum prolificum

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Abstract: A new alkaloid, named prolifine (1), was isolated along with four known alkaloids, liriodenine (2), 6-hydroxyonychine (3), isooncodine (4) and discretamine (5) from the roots of *Saccopetalum prolificum*. The structure of 1 was elucidated on the basis of spectroscopic and chemical methods.

Keywords: Saccopetalum prolificum, alkaloid, prolifine.

In recent years, an increased interest in the phytochemistry of the Annonaceae has been sparked by the isolation of the antileukemic Annonaceous acetogenin¹. *Saccopetalum prolificum* (Chun *et* How) Tsiang (Annonaceae) is an evergreen tree distributed in Hainan Province, P. R. China. The ethanolic extract of the plant exhibited cell growth inhibitory activity (20 µg/ml) against L1210 lymphocytic leukemia. A new alkaloid, prolifine (1), was isolated from the roots of *S. prolificum*, in addition to four known alkaloids, liriodenine (2), 6-hydroxyonychine (3), isooncodine (4) and discretamine (5). Among them, liriodenine (2) was reported to have potent cytotoxicity against KB, A-549, HCT-8, P-388 and L-1210 cell lines². In this paper, we focus on the structure elucidation of compound 1.

1

Prolifine (1) was isolated as yellow needles, mp 243-245°C, UV $\lambda_{\text{max}}^{\text{MeOH}}$ (log ϵ) 205 (4.43), 222 (4.27), 253 (4.02), 269 (4.06), 286 (4.04), 352 (3.97) nm. The HREIMS of **1** exhibited the [M]⁺ peak at m/z 243.0529 corresponding to the molecular formula $C_{13}H_9NO_4$ (calcd 243.0531). Its IR spectrum indicated the presence of hydroxy (3419)

cm⁻¹), carbonyl (1712 cm⁻¹) and aromatic ring (1587, 1500, 1452 cm⁻¹). The ¹³C NMR spectrum and DEPT experiments of 1 revealed 13 signals, composed of one methyl, four methines and eight quaternary carbons (Table 1). The ¹H NMR spectrum showed the existence of a methyl group at δ 2.79 (3H, s) bonded to an aromatic ring. In the low-field region of the ¹H NMR spectrum, two characteristic α- and β- pyridine protons appeared at δ 8.83 (1H, d, J = 5.2 Hz) and δ 7.59 (1H, d, J = 5.2 Hz) respectively. The ¹H NMR signal of a hydroxy group resonated at δ 13.72 due to a hydrogen bond formed between the hydroxy and the carbonyl. The presence of two vicinal hydroxy groups was demonstrated by the positive reaction of 1 to SrCl₂ test³. Then an AB pair of aromatic protons at δ 6.71 (1H, d, J = 8.8 Hz) and δ 7.04 (1H, d, J = 8.8 Hz) must be located at C-5 and C-6 positions. The ¹H and ¹³C NMR data of **1** were further assigned on the basis of the HMBC spectrum. In the HMBC spectrum of 1, the correlations of H-2 with C-3, C-4 and C-9a; H-3 with C-2, C-4, C-4a and 4-CH₃; H-5 with C-4b, C-6, C-7 and C-8a; H-6 with C-4b, C-5, C-7 and C-8; 7-OH with C-6, C-7 and C-8; 8-OH with C-7, C-8 and C-8a; the hydrogen of 4-CH₃ with C-3 and C-4 were observed. All of these correlations were in agreement with the structure.

Table 1 . ¹³ C and ¹ H NMR Spectral Data of 1 in CDCl ₃ (125 MHz for ¹³ C and 500 MHz for ¹ H)

NO.	δ_{C}	δ_{H}	$ m J_{HZ}$
2	151.0	8.83 (d)	5.2
3	126.2	7.59 (d)	5.2
4	115.4		
5	105.0	6.71 (d)	8.8
6	118.9	7.04 (d)	8.8
7	141.5		
8	146.1		
9	180.1		
4a	159.1		
4b	144.6		
8a	103.8		
9a	154.4		
7-OH		9.10 (s)	
8-OH		13.72 (s)	
CH_3	22.60	2.79 (s)	

References

- 1. F. Q. Alali, X. X. Liu, and J. L. McLaughlin, J. Nat. Prod. 1999, 62 (3), 504.
- 2. Y. C. Wu, C. Y. Duh, S. K. Wang, K. S. Chen, and T. H. Yang, J. Nat. Prod. 1990, 53 (5), 1327.
- 3. X. S. Yao, S. X. Zhao, D. J. Pan, R. Y. Zhang, M. S. Wang, Y. J. Chen, F. P. Wang, *Nat. Prod. Chem.* 2nd ed., People's Health Press, Beijing, 1997, 200.

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