Alkaloids from the Roots of Saccopetalum prolificum

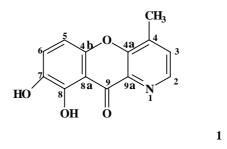
Ming Lei WANG, Jiang DU, Ruo Yun CHEN, De Quan YU*

Institute of Materia Medica, Chinese Academy of Medical Sciences and Peking Union Medical College, Beijing 100050

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 μ g/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.



Prolifine (1) was isolated as yellow needles, mp 243-245°C, UV λ_{max}^{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 C₁₃H₉NO₄ (calcd 243.0531). Its IR spectrum indicated the presence of hydroxy (3419

Ming Lei WANG et al.

cm⁻¹), carbonyl (1712 cm⁻¹) and aromatic ring (1587, 1500, 1452 cm⁻¹). The ${}^{13}C$ 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.

NO.	$\delta_{\rm C}$	$\delta_{\rm H}$	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)	

Table 1. ¹³C and ¹H NMR Spectral Data of 1 in CDCl₃ (125 MHz for ¹³C and 500 MHz for ¹H)

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.

Received 17 September 1999