Xanthones from Swertia alternifolia

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Accepted 4 August 2001 Revised 21 July 2001 Received 17 July 2000

Chemical examination of the whole plant of Swertia alternifolia yielded four xanthones, 2,8-dihydroxy-1,6-dimethoxyxanthone,1,8-dihydroxy-3,5- dimethoxyxanthone, 1,2,6,8-tetrahydroxyxanthone and 1,5,8-trihydroxy-3-methoxyxanthone. These compounds were characterised by chemical and spectral methods. These compounds were isolated for the first time from this plant.

The alcoholic extract of Swertia genus showed CNS depressant, mutagenic, antipsychotic, tuberculostatic, choleretic and antidiabetic activities1-6. Swertia chirata is a well known medicinal herb of Garhwal hills and used in Avuryedic system of medicine as laxative, febrifuge, stomachic and bitter tonic7. The chemical constituents of S.chirata responsible for its medicinal properties have been found to be mainly tetraoxygenated xanthones8. Xanthones have been evaluated for antioxidative properties and have shown to possess free radical and superoxide anion scavenging activity9-11. Due to the high demand of S.chirata and unplanned exploitation by traders, it is getting extinct in hills and people are frequently using the species S. alata and S. paniculata12, which are supposed to be equally effective. Keeping in view the commercial and pharmaceutical importance of this genus we have carried out the chemical investigation of S. alternifolia Royle¹³. Four xanthones have been isolated from the whole plant for the first time from this species.

Whole plant of *S. alternifolia* was collected from Tungnath, at an altitude of 4500 m. It is shade dried, powered (2 kg) and Soxhlet extracted with light petroleum (60-80°). The extract when concentrated under vacuum afforded compound 1 (250 mg). Petroleum ether free mass was re-extracted with ethyl acetate, concentrated *in vacuo* and after drying subjected to column chromatography over si-gel. Elution with CHCl₃:MeOH (98:2→95:5) afforded compound 2 (150 mg). Ethyl acetate-free mass was extracted with MeOH, which on

column chromatography and gradient elution with CHCl₃:MeOH (95:5→90:10) afforded compounds 3 (100 mg) and 4 (150 mg) which have been repeatedly purified by RPHPLC using CHCl₃:cyclohexane (99:1) as a solvent system.

Compound 1, yellow crystalline solid, m.p. 195° (lit. m.p¹⁴. 191-193°), molecular formula $C_{15}H_{12}O_6$, MS m/z M⁺ [288], positive to iron (III) chloride. KI exposure and 15% H_2SO_4 test, fluoresced yellow under UV light. Its UV λ_{max} MeOH 207, 230, 254, 267, 279 and 300 ¹H-NMR (C_5D_5N , TMS); δ 4.05 (3H,s lxOMe), 3.74 (3H,s, 1 x OMe), 11.8 and 13.9 (each 1H, chelated OH), 7.59 (d, J=9 Hz, H-3), 7.23 (d, J=1.5 Hz, H-7) were in accordance to 2,8-dihydroxy-1,6-dimethoxyxanthone, further confirmed by ¹³C NMR spectrum.

Compound 2, $C_{15}H_{12}O_6$, m.p. 184-185° indicated it to be a 1,3,5,8-tetraoxygenated xanthone by its UV spectrum which showed λ_{max} MeOH at 238, 260, 315 and 317 nm¹⁵. The ¹H-NMR spectrum showed the presence of two methoxyl groups at δ 3.89 and 3.96, two chelated hydroxyl groups at 11.98 and 11.39 and four aromatic protons at 6.35 and 6.54 (2H, dd, J=2.2 Hz meta protons), 7.22 and 6.71 (2H, dd, J=8.8 Hz, ortho protons). On this basis 2 was identified as 1, 8-dihydroxy-3, 5-dimethoxyxanthone¹⁵ and confirmed by ¹³C NMR spectrum.

Compound 3, yellow crystalline solid m.p. $330-331^{\circ 16}$, mol. formula $C_{13}H_8O_6$, MS m/z [M+H]* [261] gave all the chemical tests for xanthones similar to compounds 1 and 2. Its UV, IR and mass fragmentation pattern and

^{*}For correspondence

¹H-NMR chemical shifts indicated two chelated hydroxyl groups and two non-chelated hydroxyl groups with two seats of ortho and meta coupled protons, similar to 1,2.6,8-tetrahydroxyxanthone¹⁷.

Compound 4, $C_{14}H_{10}O_6$, m.p. 263-265°¹⁸ MS m/z M* [274], belonged to 1,3,5,8-tetraoxygenated series of xanthones on the basis of its UV spectrum which showed λ_{max} MeOH 201, 253, 276 and 330 nm. It was identified as 3-methoxy-1,5,8-trihydroxyxanthone on the basis of UV, IR, ¹H-NMR and MS result¹9 as well as by converting it into 1,3,5,8,-tetramethoxyxanthone, m.p. 225°¹⁶ by treating with dimethyl sulphate and potassium carbonate.

ACKNOWLEDGEMENTS

Authors are thankful to Dr. Akito Nagatsu, Faculty of Pharmaceutical Sciences, Nagoya City, University, Nagoya, 467-8603, JAPAN, for recording FAB-MS, ¹³C-NMR, DEPT and 2D-NMR spectra and All India Council for Technical Education, New Delhi for financial support.

REFERENCES

- Morimoto, I., Watnable, F., Osawa, T. and Okatsu, T., Mutat. Res.; 1982, 97, 87.
- Kanamori, H., Sakamoto, I., Mirjuta, M., Hashimoto, K. and Tanaka, O., Chem. Pharm. Bull., 1984, 32, 2290.
- Bhattacharya, S.K., Sanyal, A.K. and Ghosal, S., Bradley,
 P.B. and Dhawan, B.V. Eds., In; Drugs and Central Synaptic Transmission, Mac Millan, London, 1976, 333.
- 4. Ghosal, S. and Chaudhury, R.K., J. Pharm. Sci., 1975, 64, 88.

- 5. Ghosal. S., Biswas, K. and Chaudhury, R.K., J. Pharm. Sci., 1978, 17, 721.
- 6. Hostettman, K. and Wagner, H., Phytochemistry, 1977, 16, 821.
- Chopra, R.N., Nayar, S.L. and Chopra, S.C., In; Glossary of Indian Medicinal Plants, C.S.I.R., New Delhi, 1956, 237.
- 8. Chakravarty, A.K., Mukhopadhyay, S., Masuda, K. and Ageta, H., Indian J. Chemistry, 1992, 31B, 70.
- Minami, H., Kinoshita, M., Fukuyama, Y., Kodama, M., Yoshizawa, T., Sugiura, M., Nakagawa, K. and Tago, H., Phytochemistry, 1994, 36, 501.
- Chang, C.H., Lin, C.C., Hattori, M. and Namba, T., J. Ethnopharmacol. 1994, 44, 79.
- 11. Minami, H., Kuwayama, A., Yoshizawa, T. and Fukuyama; Y., Chem. Pharm. Bull. 1996, 44, 2103.
- Khetwal, K.S. and Verma, D.L., Indian J. Pharm. Sci., 1984, 46, 25.
- Royle, J.F. In; Illustrations of the Botany and other branches of the Natural History of the Himalayan Mountains and of the Flora of Cashmere. W.N. Allens & Co, London, 1833, 1840.
- 14. Rivialle, P., Massicot, J., Guyot, M. and Pourier, V., Phytochemistry, 1969, 8, 1533.
- Stout, G.H., Christensen, E.N., Balkenhol, W.J. and Stevens, K.L., Tetrahedron, 1969, 25, 1961.
- 16. Neshta, N.M., Glyzin, V.J., Nikolaeva, G.C. and Sheichenko, V.I., Khim. Prir. Soedin, 1980, 106
- Carpenter, T., Locksley, H.D. and Scheinmann, F., Phytochemistry, 1969, 8, 2013.
- 18. Ghosal, S., Sharma, P.V., Chaudhury. R.K. and Bhattacharya, S.K., J. Pharm. Sci., 1973, 62, 926.
- 19. Kanamori, H., Sakamoto, J., Mirjuta, M., Hashimoto. K. and Tanaka. O., Chem. Pharm. Bull., 1984, 32, 2290.