A Hypnotic Barbitone from Sphaeroides oblongus

S. K. Mitra, B. Sanyal, S. N. Ganguly, and Biswapati Mukherjee*a

^a Department of Pharmacology, B.C. Roy Post-graduate Institute of Basic Medical Sciences, 244B Acharya J. C. Bose Road, Calcutta 700020, India

b Bose Institute, Calcutta 700009, India

The common hypnotic 5,5-diethylbarbituric acid has been isolated for the first time from a natural source, *Sphaeroides oblongus*.

The puffer fish *Sphaeroides oblongus* is responsible for occasional damage¹ to health and lives in the coastal region of India. A chemical investigation of the major source of poison (the liver²) of *S. oblongus*, collected from the coast of the Bay of Bengal, afforded shining white needles, m.p. 195—196 °C. Spectroscopic data [λ_{max} 224(EtOH); 224 and 240(EtOH-NaOH); and 243 nm (0.1 n-NaOH); ν_{max} (KBr) 1682s, 1721s, and 1769s (3 × C=0), 3430(br, 3210m, 3170m, and 3080s cm⁻¹ (highly associated NH); δ_{H} 0.7 (3H, t, CH₂CH₃) and 1.8 (2H, q, CH₂CH₃); m/z 156 (M^+ , C₆H₈N₂O₃), 155, 141, 112, and 98] indicated a structure similar to 5-ethylbarbituric acid (1), but comparison (mixed m.p., t.l.c., i.r.) with the synthetic product³ showed them to be different. ¹³C N.m.r. signals of the isolated compound [δ 9.08 (q, CH₃), 31.11 (t, CH₂), 56.44 (s, quat. C), 149.91 (s, C=O), and 173.04 (s, 2 × C=O)] were

similar to those of structure (1) excepting for the peak at δ 56.44. The molecular weight (M^+ 184) and composition ($C_8H_{12}N_2O_3$) of the compound were determined by fast atomic bombardment mass spectrometry leading to its identification as 5,5-diethylbarbituric acid (barbitone) (2); this was confirmed by spectroscopic comparison with authentic material.

The occurrence of barbitone in *S. oblongus* was monitored for two years; its appearance was found to be seasonal, probably related to the breeding term.

Tetrodotoxin (3) is the major constituent of the puffer fish elsewhere.⁴ The structural patterns of tetrodotoxin (3) and barbitone (2) show some similarity.

We thank Professor Maurice Shamma and Mr. Alan J. Freyer, Pennsylvania State University, U.S.A., for the spectra.

Received, 19th January 1988; Com. 8/00341F

References

1 S. Jones, Ind. J. Med. Res., 1956, 44, 353.

2 C. Y. Kao, Pharmacol. Rev., 1966, 18, 997.

3 J. Shapira, J. Org. Chem., 1962, 27, 1918.

4 T. Goto, Y. Kishi, S. Takahashi, and Y. Hirata, *Tetrahedron*, 1965, 21, 2059.