―天然薬物ノート―

## A Morphinane Alkaloid, Sinococuline, from Stereospermum suaveolens DC.

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A morphinane alkaloid, sinococuline was isolated from the root of *Stereospermum* suaveolens DC (Bignoniaceae) and showed cytotoxic activity. The structure was elucidated on the basis of spectroscopic data. This is the first report of its isolation from this plant.

Keywords Sinococuline; Stereospermum suaveolens DC; Bignoniaceae; morphinane alkaloid

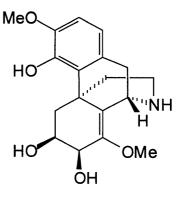
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Stereospermum suaveolens DC, a member of the Bignoniaceae family, is distributed throughout India (Indian name Padal, Patala). The root of this plant has been used in India traditional medicine for anti-fever purposes. From this plant, lapachol, flavone glycosides,  $\beta$ -sitosterol, anthraquinone and naphtoquinone <sup>1)-5)</sup> have previously been reported.

In this paper, we report the bioassay-guided isolation of sinococuline, a rare morphinane alkaloid, from the root of *S. suaveolens* DC.

The roots of S. suaveolens were purchased from Associated Dichem Corporation (India). Dried roots of S. suaveolens (1080 g) were extracted with MeOH at room temperature for 1 week. The MeOH solution was filtered and the filtrate was concentrated in vacuo to yield the MeOH extract (23 g). The MeOH extract (10.2 g) was adsorbed on silica gel and applied to column chromatography over silica gel and eluted with hexane-EtOAc (4 : 1, v/v), EtOAc and MeOH to afford three fractions, fraction 1 (519 mg), fraction 2 (500 mg) and fraction 3 (8134 mg) and the cytotoxic activity was detected in fraction 3. A portion of fraction 3 (400 mg) was subjected to gel filtration on a Sephadex LH-20 column (11 mm id x 950 mm). Elution with MeOH gave fraction 3-1 (30 mg), 3-2 (186 mg) and 3-3 (189 mg) (Kd value=0 to 0.75, 0.75 to 1.0 and 1.0 to 2.0, respectively). The active fraction, 3-2, was further purified by silica gel column chromatography using CHCl<sub>3</sub>-MeOH (9 : 1, v/v) as an eluent, followed by Sephadex LH-20 (MeOH) to yield compound 1 (74 mg).

Compound 1 was obtained as a white powder. The molecular formula was determined to be  $C_{18}H_{23}NO_5$  by high-resolution electro-spray-ionization mass spectra (HR-ESI-MS). When the NMR data for 1 were compared with those of a known morphinane alkaloid, sinococuline, isolated from *Cocculus trilobus*, <sup>6)</sup> a similar chemical shift pattern was observed. Detailed comparison of its spectroscopic data ( $[\alpha]_D$ , MS, <sup>1</sup>H- and <sup>13</sup>C-NMR) with those reported in the literature<sup>6), 7)</sup> identified the structure of the compound 1 as sinococuline.



Sinococuline (1)

Sinococuline showed cytotoxic activity against the human lung cancer cell line A549 ( $IC_{50} = 1.83 \mu M$ ) and colorectal cancer cell line HT-29 ( $IC_{50} = 2.08 \mu M$ ).

Phytochemical studies on the genus *Stereospermum* have found lignans, anthraquinones and naphthoquinones.<sup>8)-13)</sup> No morphinane alkaloids from

Stereospermum genus have been investigated. Sinococuline is a rare morphinane alkaloid previously reported from only Menispermaceae family sources, *C. trilobus*,<sup>6)</sup> Stephania cepharantha,<sup>7)</sup> S. excentrica,<sup>14)</sup> S. *sutchuenensis* H.S.Lo<sup>15)</sup> and Strychnopsis thouarsii Baillon.<sup>16)</sup> This is the first report on the isolation of sinococuline from plants of the Bignoniaceae family.

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