

## Fissohamione, a Novel Furanone from Fissistigma oldhamii

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Abstract: Fissohamione (1), a novel (R)-4,5-dimethoxy-3-(4'-phenyl-2'-oxobutyl)-5H-furan-2-one, has been isolated from Fissistigma oldhamii. The structure of 1 was elucidated by spectroscopic methods. © 1999 Elsevier Science Ltd. All rights reserved.

Our laboratory and other investigators have reported a series of studies on *Fissistigma* species.<sup>1-10</sup> *Fissistigma oldhamii* (Memsl.) Merr. (Annonaceae) is a perennial shrub, which is distributed mainly in southern China and Taiwan. This plant has been used as a folklore medicine for liver protection, anti-inflammatory and anti-arthritic effects, and anti-tumor action.<sup>8</sup> As part of our continuing investigation on the phytochemical and bioactive principles of Formosan Annonaceous plants, we have isolated fissohamione (1), a novel 3,4,5-substituted furan-2-one, from the methanolic extract of the seeds of *F. oldhamii*.

Specimens of *F. oldhamii* were collected from Taichung, Taiwan in August 1989. The methanolic extract was separated by column chromatography using 1:2 EtOAc/Hexane as an eluent to obtain a fraction, which was further purified by preparative TLC to yield fissohamione (1, 0.003% dry wt.).

Fissohamione (1) was obtained as a pale-yellow oil,  $[\alpha]_D$ = +4.6 (c 0.25, CHCl<sub>3</sub>). The molecular formula,  $C_{16}H_{18}O_5$ , was confirmed by high-resolution mass spectroscopic measurement (m/z 290.1157 [M]<sup>+</sup>, calcd. 290.1154). The presence of an  $\alpha$ , $\beta$ -unsaturated lactone and ketone moieties were substantiated by its UV ( $\lambda$  max at 210 and 235 nm) and IR ( $\nu_{max}$  at 1765, 1721 and 1681 cm<sup>-1</sup>) spectra. The <sup>1</sup>H NMR spectrum of 1 (Table 1) indicated the presence of signals corresponding to one mono-substituted aromatic ring, one proton, two methoxyls, one methylene and two methylene groups. The <sup>13</sup>C NMR and DEPT experiments (Table 1) further confirmed the possession of six methines, five quaternary carbons (including a carbonyl at  $\delta$  204.91), three methylenes and two methoxyl signals. The above mentioned data indicated the presence of a 4'-phenyl-2'-oxobutyl side chain, which is in agreement with those reported in the literature. <sup>10</sup>

Further confirmation of the structural assignment, two-dimensional NMR experiments were employed. The sequential correlations of NOESY and long range HETCOR were successfully established as shown in Figure 1 and Table 1, respectively. The absolute configuration of 1 was determined by the CD method. According to a negative  $\pi$ - $\pi$ \* Cotton effect ( $\Delta \varepsilon < 0$ ), it clearly indicated the  $\gamma$ -butenolide fragment has the (R)-configuration. The above results support the structure of 1 as a novel (R)-4,5-dimethoxy-3-(4'-phenyl-2'-oxobutyl)-5H-furan-2-one, which is provisionally named fissohamione.

Fig. 1 NOESY experiments of fissohamione (1)

Table 1. <sup>13</sup>C NMR (100 MHz, CDCl<sub>3</sub>) and <sup>1</sup>H NMR (400 MHz) data for fissohamione (1).

Position	δC	δH, J (Hz)	LRHETCOR
2	171.26		
3	98.85		
4	170.30		
5	97.91	5.68, s	C-4
1'	36.35	3.37, ABq (17.6)	C-2', C-2, C-4, C-3
2'	204.91		
3'	43.91	2.84, m	
4'	29.62	2.91, m	
5'	140.60		
6',10'	128.3 🖺		
7',9'	128.48	7.16-7.27, m	
8'	لـ126.1 <i>6</i>		
OMe-4	58.35	3.93, s	C-4
OMe-5	55.28	3.52, s	C-5

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