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## LIGNANS AND NEOLIGNAN FROM THE STEMS OF *CANSJERA RHEEDEI* J.F.GMEL.

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### Summary

*Cansjera rheedei* J.F.Gmel. belongs to the family Opiliaceae, which is distributed in the regions of southern Asia. Research on this plant's chemical constituents has been limited to the identification of several common phenolic compounds. This phytochemical study was conducted on the stems of *C. rheedei* to isolate five lignans and one neolignan. The chemical structures of six isolated compounds were elucidated by using spectral data analyses and comparison with those in references, leading to the determination of alyterinate A (**1**), alyterinate B (**2**), (+)-pinoresinol (**3**), (+)-medioresinol (**4**), (+)-syringaresinol (**5**), and balanophonin (**6**). All of them were reported for the first time from the genus *Cansjera*.

**Keywords:** *Cansjera rheedei*; Lignan; Neolignan.

### 1. Introduction

*Cansjera* is a small genus in the family Opiliaceae that consists of three species (*C. rheedei*, *C. leptostachya*, and *C. parvifolia*). *Cansjera rheedei* J.F.Gmel., a shrub or climbing shrub, is distributed from India and Nepal to southern China (Vietnam, Thailand, Laos, and Myanmar) and western Malaysia [1]. The ethanol extract of the aerial parts of *C. rheedei* showed a nontoxic effect through the evaluation of acute and sub-acute toxicity in animal trials [2]. Several investigations into the biological activities of this herb were performed to illustrate anthelmintic [3], hepatoprotective [4], hypoglycemic [5], and nephroprotective [6] effects. To date, only a single study has investigated the phytochemistry of *C. rheedei*, reporting several common phenolic and flavonoid compounds, including caffeic acid, ferulic acid, quercetin, quercetin-3-*O*- $\beta$ -D-glucoside, and rutin [7]. Here, we report the isolation and structural elucidation of six compounds from the stems of *C. rheedei*, consisting of five lignans (alyterinate A (**1**), alyterinate B (**2**), (+)-pinoresinol (**3**), (+)-medioresinol (**4**), and (+)-syringaresinol (**5**)) and one neolignan (balanophonin (**6**)). All these chemical constituents were reported for the first time from this genus.

### 2. Materials and methods

#### 2.1. Plant material

Stems of *C. rheedei* J.F.Gmel. were collected from Bac Giang province (June 2024) and authenticated by Dr. Vo Thanh Hoa (Department of Pharmacognosy, University of Health Sciences, Vietnam National University, Ho Chi Minh City). A voucher specimen of this sample

(CRS2024-01) was deposited in the Department of Pharmacognosy, University of Health Sciences, Vietnam National University, Ho Chi Minh City.

#### 2.2. General experimental procedures

Column chromatography was performed with *silica gel* (60  $\mu$ m, Chromatorex MB70-40/75, Japan) and Sephadex LH-20 (GE Healthcare, Sweden). Thin-layer chromatography (TLC) was conducted on *silica gel* 60 F<sub>254</sub> plates (Merck, Germany). The reversed-phase preparative HPLC was performed using an LC-40D instrument (Shimadzu, Japan) connected to a Cosmosil C<sub>18</sub>-AR-II column (5  $\mu$ m, 20 mm I.D.  $\times$  250 mm) with a UV detector. Optical rotations were recorded by a digital polarimeter (Jasco P-2000, Japan). HR-ESIMS data were obtained with the Shimadzu LCMS-9030 series Q-TOF mass spectrometer analyzer (Shimadzu, Japan). 1D and 2D NMR spectra were measured on a Bruker AVIII600 spectrometer (Bruker, Germany) using TMS as an internal standard. All reagents and solvents were of analytical grade.

#### 2.3. Extraction and isolation

The plant samples were air-dried and ground to a raw powder before further processing. The dried powdered stems of *C. rheedei* (3.5 kg) were extracted with 95% ethanol (35 L, 3 times) at 50°C. The solvent was removed under reduced pressure to give the crude extract, which was then dispersed in distilled water (2 L) and partitioned with ethyl acetate (4 times, each 2 L) and continuously with *n*-butanol (3 times, each 2 L). The organic solvents were evaporated *in vacuum* to yield ethyl acetate extract (CR-A, 62 g) and *n*-butanol extract (CR-B, 40 g).



The CR-A extract was separated by *silica gel* column chromatography with elution of a gradient solvent mixture of *n*-hexane - ethyl acetate (9:1 to 0:10, v/v) to obtain 14 fractions (A.1 - A.14). The fraction A.7 was purified by preparative HPLC (40 % acetonitrile (ACN) in water, isocratic, 6 mL/min) to afford compounds **1** (32.6 mg,  $t_R$  16.2 min) and **2** (14.1 mg,  $t_R$  18.5 min). The fraction A.9 was chromatographed by *silica gel* column with gradient elution of chloroform - ethyl acetate (9:1 - 5:5, v/v) and applied subsequently on Sephadex LH-20 column to furnish compounds **3** (45.0 mg) and **4** (148.4 mg). The fraction A.10 was purified by preparative HPLC (25 % ACN in water, isocratic, 6 mL/min) to obtain compounds **5** (81.6 mg,  $t_R$  12.5 min) and **6** (4.5 mg,  $t_R$  14.8 min).

**Compound 1 (alyterinate A):** colorless needle crystals;  $[\alpha]_D^{25} + 12.6$  ( $c$  0.1, MeOH); HR-ESIMS (+)  $m/z$  371.1490  $[M-H_2O+H]^+$ ;  $^1H$ -NMR (600 MHz,  $CD_3OD$ )  $\delta_H$  (ppm): 6.90 (1H, d,  $J = 1.8$  Hz, H-2), 6.75 (1H, m, H-5), 6.76 (1H, m, H-6), 5.15 (1H, d,  $J = 7.2$  Hz, H-7), 3.11 (1H, dd,  $J = 8.4, 7.2$  Hz, H-8), 6.76 (1H, m, H-2'), 6.71 (1H, d,  $J = 7.8$  Hz, H-5'), 6.61 (1H, dd,  $J = 7.8, 1.8$  Hz, H-6'), 2.74 (1H, dd,  $J = 13.8, 6.0$  Hz, H-7'a), 2.54 (1H, dd,  $J = 13.8, 10.2$  Hz, H-7'b), 2.97 (1H, m, H-8'), 4.07 (1H, dd,  $J = 8.4, 6.6$  Hz, H-9'a), 3.76 (1H, dd,  $J = 8.4, 6.6$  Hz, H-9'b), 3.65 (3H, s, 9-OCH<sub>3</sub>), 3.83 (6H, s, 3/3'-OCH<sub>3</sub>);  $^{13}C$ -NMR (150 MHz,  $CD_3OD$ )  $\delta_C$  (ppm): 134.2 (C-1), 110.6 (C-2), 149.0 (C-3), 147.4 (C-4), 116.1 (C-5), 119.8 (C-6), 84.0 (C-7), 57.1 (C-8), 174.0 (C-9), 132.3 (C-1'), 113.4 (C-2'), 149.0 (C-3'), 146.1 (C-4'), 116.2 (C-5'), 122.2 (C-6'), 35.4 (C-7'), 45.3 (C-8'), 74.0 (C-9'), 52.2 (9-OCH<sub>3</sub>), 56.4 (3-OCH<sub>3</sub>), 56.4 (3'-OCH<sub>3</sub>).

**Compound 2 (alyterinate B):** colorless needle crystals;  $[\alpha]_D^{25} + 15.1$  ( $c$  0.1, MeOH); HR-ESIMS (+)  $m/z$  387.1435  $[M+H]^+$ , 369.1339  $[M-H_2O+H]^+$ ;  $^1H$ -NMR (600 MHz,  $CD_3OD$ )  $\delta_H$  (ppm): 6.95 (1H, d,  $J = 1.8$  Hz, H-2), 6.77 (1H, d,  $J = 7.8$  Hz, H-5), 6.80 (1H, m, H-6), 5.12 (1H, d,  $J = 7.8$  Hz, H-7), 3.70 (1H, dt,  $J = 7.8, 2.4$  Hz, H-8), 6.75 (1H, d,  $J = 1.8, H-2'$ ), 6.79 (1H, m, H-5'), 6.65 (1H, dd,  $J = 8.4, 1.8$  Hz, H-6'), 6.43 (1H, q,  $J = 2.4$  Hz, H-7'), 4.89 (1H, dd,  $J = 13.8, 3.0$  Hz, H-9'a), 4.75 (1H, dt,  $J = 13.8, 3.0$  Hz, H-9'b), 3.83 (3H, s, 3-OCH<sub>3</sub>), 3.77 (3H, s, 9-OCH<sub>3</sub>), 3.86 (3H, s, 3'-OCH<sub>3</sub>);  $^{13}C$ -NMR (150 MHz,  $CD_3OD$ )  $\delta_C$  (ppm): 132.5 (C-1), 110.7 (C-2), 149.1 (C-3), 147.7 (C-4), 116.1 (C-5), 120.2 (C-6), 84.1 (C-7), 59.8 (C-8), 173.5 (C-9), 130.1 (C-1'), 113.1 (C-2'), 149.0 (C-3'), 147.3 (C-4'), 116.4 (C-5'), 122.5 (C-6'), 124.3 (C-7'), 137.7 (C-8'), 71.2 (C-9'), 52.8 (9-OCH<sub>3</sub>), 56.4 (3-OCH<sub>3</sub>), 56.4 (3'-OCH<sub>3</sub>).

**Compound 3 ((+)-pinoresinol):** colorless needle crystals;  $[\alpha]_D^{25} + 20.2$  ( $c$  0.1, MeOH); HR-ESIMS (+)  $m/z$  359.1488  $[M+H]^+$ , 341.1376  $[M-H_2O+H]^+$ ;  $^1H$ -NMR (600 MHz,  $CD_3OD$ )  $\delta_H$  (ppm): 6.93 (2H, d,  $J = 1.8$  Hz, H-2/2'), 6.77 (2H, d,  $J = 8.4$  Hz, H-5/5'), 6.79 (2H, dd,  $J = 8.4, 1.6$  Hz, H-6/6'), 4.68 (2H, d,  $J = 4.2$  Hz, H-7/7'), 3.10 (2H, q,  $J = 4.2$  Hz, H-8/8'), 4.20 (2H, dd,  $J = 9.0, 6.6$  Hz, H-9a/9'a), 3.81 (2H, dd,  $J = 9.0, 3.6$  Hz, H-9b/9'b), 3.83 (6H, s, 3/3'-OCH<sub>3</sub>);  $^{13}C$ -NMR (150 MHz,  $CD_3OD$ )  $\delta_C$  (ppm): 133.8 (C-1/1'), 111.0 (C-2/2'), 149.1 (C-3/3'), 147.2 (C-4/4'), 116.1 (C-5/5'), 120.0 (C-6/6'), 87.4 (C-7/7'), 55.3 (C-8/8'), 72.6 (C-9/9'), 56.4 (3/3'-OCH<sub>3</sub>).

**Compound 4 ((+)-medioresinol):** amorphous white powder;  $[\alpha]_D^{25} + 18.5$  ( $c$  0.1, MeOH); HR-ESIMS (+)  $m/z$  389.1593  $[M+H]^+$ , 371.1494  $[M-H_2O+H]^+$ ;  $^1H$ -NMR (600 MHz,  $CD_3OD$ )  $\delta_H$  (ppm): 6.66 (2H, s, H-2/6), 4.71 (1H, d,  $J = 4.2$  Hz, H-7), 3.14 (1H, m, H-8), 4.25 (1H, m, H-9a), 3.88 (1H, m, H-9b), 6.95 (1H, d,  $J = 1.8$  Hz, H-2'), 6.77 (1H, d,  $J = 7.8$  Hz, H-5'), 6.81 (1H, dd,  $J = 7.8, 1.8$  Hz, H-6'), 4.71 (1H, d,  $J = 4.2$  Hz, H-7'), 3.14 (1H, m, H-8'), 4.25 (1H, m, H-9'a), 3.88 (1H, m, H-9'b), 3.85 (6H, s, 3/5-OCH<sub>3</sub>), 3.86 (3H, s, 3'-OCH<sub>3</sub>);  $^{13}C$ -NMR (150 MHz,  $CD_3OD$ )  $\delta_C$  (ppm): 133.2 (C-1), 104.5 (C-2/6), 149.4 (C-3/5), 136.2 (C-4), 87.7 (C-7), 55.6 (C-8), 72.7 (C-9), 133.8 (C-1'), 111.0 (C-2'), 149.1 (C-3'), 147.3 (C-4'), 116.1 (C-5'), 120.1 (C-6'), 87.5 (C-7'), 55.3 (C-8'), 72.6 (C-9'), 56.8 (3/5-OCH<sub>3</sub>), 56.4 (3'-OCH<sub>3</sub>).

**Compound 5 ((+)-syringaresinol):** colorless needle crystals;  $[\alpha]_D^{25} + 23.9$  ( $c$  0.1, MeOH); HR-ESIMS (+)  $m/z$  419.1701  $[M+H]^+$ , 401.1602  $[M-H_2O+H]^+$ ;  $^1H$ -NMR (600 MHz,  $CD_3OD$ )  $\delta_H$  (ppm): 6.65 (4H, s, H-2/6/2'/6'), 4.71 (2H, d,  $J = 4.2$  Hz, H-7/7'), 3.13 (2H, m, H-8/8'), 4.26 (2H, m, H-9a/9'a), 3.87 (2H, dd,  $J = 9.0, 3.6$  Hz, H-9b/9'b), 3.84 (12H, s, 3/5/3'/5'-OCH<sub>3</sub>);  $^{13}C$ -NMR (150 MHz,  $CD_3OD$ )  $\delta_C$  (ppm): 133.1 (C-1/1'), 104.5 (C-2/6/2'/6'), 149.3 (C-3/5/3'/5'), 136.2 (C-4/4'), 87.6 (C-7/7'), 55.5 (C-8/8'), 72.8 (C-9/9'), 56.8 (3/5/3'/5'-OCH<sub>3</sub>).

**Compound 6 (balanophonin):** amorphous white powder;  $[\alpha]_D^{25} + 10.0$  ( $c$  0.1, MeOH); HR-ESIMS (+)  $m/z$  357.1330  $[M+H]^+$ , 339.1228  $[M-H_2O+H]^+$ ;  $^1H$ -NMR (600 MHz,  $CD_3OD$ )  $\delta_H$  (ppm): 6.95 (1H, d,  $J = 1.8$  Hz, H-2), 6.78 (1H, d,  $J = 8.4$  Hz, H-5), 6.83 (1H, dd,  $J = 8.4, 1.8$  Hz, H-6), 5.61 (1H, d,  $J = 6.6$  Hz, H-7), 3.57 (1H, q,  $J = 6.0$  Hz, H-8), 3.85 (2H, m, H-9), 7.24 (1H, d,  $J = 1.8$  Hz, H-2'), 7.29 (1H, brs, H-6'), 7.63 (1H, d,  $J = 15.6$  Hz, H-7'), 6.69 (1H, dd,  $J = 15.6, 7.8$  Hz, H-8'), 9.59 (1H, d,  $J = 7.8$  Hz, H-9'), 3.82 (3H, s, 3-OCH<sub>3</sub>), 3.92 (3H, s, 3'-OCH<sub>3</sub>);  $^{13}C$ -NMR (150 MHz,  $CD_3OD$ )  $\delta_C$  (ppm): 133.9 (C-1), 110.6 (C-2), 149.2 (C-3), 147.8 (C-4), 116.2 (C-

5), 119.8 (C-6), 90.1 (C-7), 54.7 (C-8), 64.6 (C-9), 129.6 (C-1'), 114.3 (C-2'), 146.0 (C-3'), 153.0 (C-4'), 131.3 (C-5'), 120.0 (C-6'), 156.1 (C-7'), 127.1 (C-8'), 196.1 (C-9'), 56.4 (3-OCH<sub>3</sub>), 56.8 (3'-OCH<sub>3</sub>).

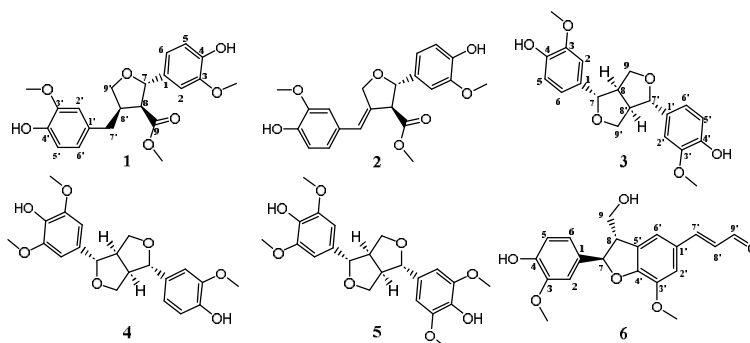


Fig.1. Chemical structures of the isolated compounds (1–6)

### 3. Results and discussion

The 95% ethanol extract from the stems of *C. rheedei* was partitioned and purified using various chromatographic methods to furnish six compounds (1–6). The chemical structures of the isolated compounds were determined by their spectral data and comparison with those in the corresponding references.

Compound **1** was obtained as colorless needle crystals. The molecular formula was identified to be C<sub>21</sub>H<sub>24</sub>O<sub>7</sub> by its HR-ESIMS data at  $m/z$  371.1490 [M-H<sub>2</sub>O+H]<sup>+</sup> (calcd. for C<sub>21</sub>H<sub>23</sub>O<sub>6</sub><sup>+</sup>, 371.1495). The <sup>1</sup>H-NMR spectrum of **1** showed six protons of two ABX systems at the olefinic region (6.90 - 6.61 ppm); three methine proton signals at  $\delta_H$  5.15 (1H, d, 7.2 Hz), 3.11 (1H, dd, 8.4, 7.2 Hz), and 2.97 (1H, m); two methylene in the range of 4.07 - 2.54 ppm; and three methoxy groups at  $\delta_H$  3.83 (6H, s) and 3.65 (3H, s). The <sup>13</sup>C-NMR spectrum of **1** illustrated 21 signals, including one carbonyl carbon at  $\delta_C$  174.0; four oxygenated olefinic carbons ( $\delta_C$  149.0 - 146.1); two oxygenated methine and methylene groups at  $\delta_C$  84.0 and 74.0, respectively; and three methoxy signals ( $\delta_C$  56.4, 56.4, and 52.2). The proton at  $\delta_H$  3.11 (H-8) exhibited the COSY correlation with  $\delta_H$  2.97 (H-8') and the HMBC interaction to  $\delta_C$  74.0 (C-9') proved that compound **1** was a furan lignan. In the HMBC spectrum, correlations from the methylene proton ( $\delta_H$  2.74 and 2.54, H-7') to  $\delta_C$  57.1 (C-8) and 74.0 (C-9') were observed. Furthermore, three methoxy protons interacted with two aromatic carbons at  $\delta_C$  149.0 and the carbonyl group at  $\delta_C$  174.0. Spectral data analyses and optical rotation  $[\alpha]_D^{25} + 12.6$  led to the elucidation of compound **1** as alyterinate A, which showed similarity with those of the published literature [8].

Compound **2** was obtained as colorless needle crystals. The molecular formula was identified to

be C<sub>21</sub>H<sub>22</sub>O<sub>7</sub> by its HR-ESIMS data at  $m/z$  387.1435 [M+H]<sup>+</sup> (calcd. for C<sub>21</sub>H<sub>23</sub>O<sub>7</sub><sup>+</sup>, 387.1438) and  $m/z$  369.1339 [M-H<sub>2</sub>O+H]<sup>+</sup> (calcd. for C<sub>21</sub>H<sub>21</sub>O<sub>6</sub><sup>+</sup>, 369.1338). The <sup>1</sup>H-NMR spectrum of **2** showed six protons of two ABX systems at the olefinic region (6.95 - 6.65 ppm); another olefinic proton at  $\delta_H$  6.43 (1H, q, 2.4 Hz); two methine protons at  $\delta_H$  5.12 (1H, d, 7.8 Hz) and 3.70 (1H, dt, 7.8, 2.4 Hz); and three methoxy groups at  $\delta_H$  3.86 (3H, s), 3.83 (3H, s) and 3.77 (3H, s). The <sup>13</sup>C-NMR spectrum of **2** demonstrated 21 carbons with one carbonyl signal at  $\delta_C$  173.5; four hydroxyl aromatic carbons ( $\delta_C$  149.0 - 147.3); one oxygenated methine at  $\delta_C$  84.1; one oxygenated methylene at  $\delta_C$  71.2; and three methoxy groups ( $\delta_C$  56.4, 56.4, and 52.8). In the HMBC spectrum, the proton at  $\delta_H$  5.12 (H-7) showed the correlations with  $\delta_C$  137.7 (C-8') and 71.2 (C-9'). The structure of compound **2** was a furan lignan similar to compound **1**. In addition, proton  $\delta_H$  6.43 (H-7') correlated with  $\delta_C$  71.2 (C-9') and 59.8 (C-8), indicating the appearance of one double bond at the C-7' and C-8' position. Carbon chemical shift values of C-7' ( $\delta_C$  124.3) and C-8' ( $\delta_C$  137.7) were consistent with literature values for *Z*-configured C-7'/C-8' [8], suggesting the *Z* geometry of this double bond. This compound showed  $[\alpha]_D^{25} + 15.1$  ( $c$  0.1, MeOH). Based on the spectral analyses and comparison with the reference [8], compound **2** was identified as alyterinate B.

Compound **3** was obtained as colorless needle crystals. The molecular formula was identified to be C<sub>20</sub>H<sub>22</sub>O<sub>6</sub> by its HR-ESIMS data at  $m/z$  359.1488 [M+H]<sup>+</sup> (calcd. for C<sub>20</sub>H<sub>23</sub>O<sub>6</sub><sup>+</sup>, 359.1489) and  $m/z$  341.1376 [M-H<sub>2</sub>O+H]<sup>+</sup> (calcd. for C<sub>20</sub>H<sub>21</sub>O<sub>5</sub><sup>+</sup>, 341.1389). The <sup>1</sup>H-NMR spectrum of **3** showed three protons of an ABX system (6.93 - 6.77 ppm); two methine protons at  $\delta_H$  4.68

(1H, d, 4.2 Hz) and 3.10 (1H, q, 4.2 Hz); a methylene group at  $\delta_H$  4.20 (1H, dd, 9.0, 6.6 Hz) and 3.81 (1H, dd, 9.0, 3.6 Hz); and a methoxy group at  $\delta_H$  3.83 (3H, s). The  $^{13}\text{C}$ -NMR spectrum of **3** exhibited 10 signals of a symmetrical structure, including two olefinic carbons connected with oxygen ( $\delta_C$  149.1 and 147.2); the oxygenated methine and methylene carbons at  $\delta_C$  87.4 and 72.6, respectively; and one methoxy group at  $\delta_C$  55.3. In the HMBC spectrum, proton  $\delta_H$  4.68 interacted with  $\delta_C$  72.6 and 55.3. In addition, the proton  $\delta_H$  3.10 had COSY correlations with the methine proton ( $\delta_H$  4.68) and the methylene group ( $\delta_H$  4.20 and 3.81) featured for a furofuran lignan derivative. The optical rotation of compound **3** was recorded as  $[\alpha]_D^{25} + 20.2$  ( $c$  0.1, MeOH). In comparison with spectral data in the published literature [9], compound **3** was identified as (+)-pinoresinol.

Compound **4** was obtained as an amorphous white powder. The molecular formula was identified to be  $\text{C}_{21}\text{H}_{24}\text{O}_7$  by its HR-ESIMS data at  $m/z$  389.1593  $[\text{M}+\text{H}]^+$  (calcd. for  $\text{C}_{21}\text{H}_{25}\text{O}_7^+$ , 389.1595) and  $m/z$  371.1494  $[\text{M}-\text{H}_2\text{O}+\text{H}]^+$  (calcd. for  $\text{C}_{21}\text{H}_{23}\text{O}_6^+$ , 371.1495). The  $^1\text{H}$ -NMR spectrum of **4** showed three protons of an ABX system (6.95 - 6.77 ppm); two protons of the symmetrical trisubstituted aromatic ring at  $\delta_H$  6.66 (2H, s); two oxygenated methine protons at  $\delta_H$  4.71 (2H, d, 4.2 Hz); two oxygenated methylene groups at  $\delta_H$  4.25 (2H, m) and 3.88 (2H, m); and three methoxy groups at  $\delta_H$  3.86 (3H, s), and 3.85 (6H, s). The  $^{13}\text{C}$ -NMR spectrum of **4** declared 18 signals consisting of 4 aromatic carbons linked with oxygen (149.4 - 136.2 ppm); two oxygenated methine at  $\delta_C$  87.7 and 87.5; two oxygenated methylene at  $\delta_C$  72.7 and 72.6; and three methoxy groups at  $\delta_C$  56.8 and 54.4. The NMR spectral data of compound **4** showed similar characteristics to those of compound **3**, suggesting a furofuran lignan derivative. The difference was the addition of a methoxy group to create the symmetrical 1,3,4,5-substituted phenolic unit. In addition, compound **4** indicated the optical rotation as  $[\alpha]_D^{25} + 18.5$  ( $c$  0.1, MeOH). The structure of compound **4** was elucidated as (+)-medioresinol, which was confirmed by reference [10].

Compound **5** was obtained as colorless needle crystals. The molecular formula was identified to be  $\text{C}_{22}\text{H}_{26}\text{O}_8$  by its HR-ESIMS data at  $m/z$  419.1701  $[\text{M}+\text{H}]^+$  (calcd. for  $\text{C}_{22}\text{H}_{27}\text{O}_8^+$ , 419.1700) and  $m/z$  401.1602  $[\text{M}-\text{H}_2\text{O}+\text{H}]^+$  (calcd. for  $\text{C}_{22}\text{H}_{25}\text{O}_7^+$ , 401.1600). The  $^1\text{H}$ -NMR and  $^{13}\text{C}$ -NMR spectra of **5** showed the characteristic of a furofuran lignan with a

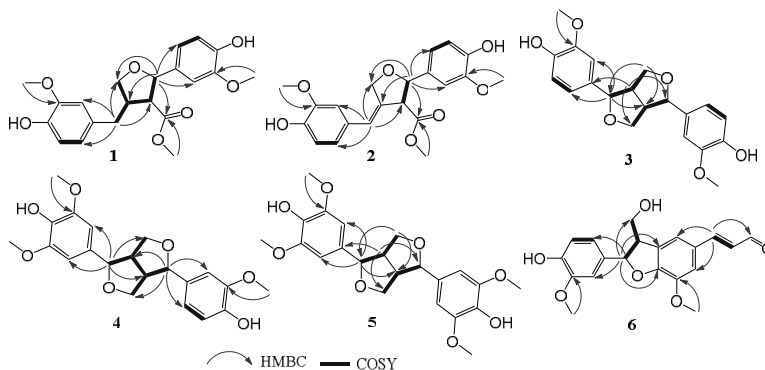
symmetrical structure like compound **3**. Furthermore, two aromatic ring units were all substituted by two methoxy groups ( $\delta_C$  56.8,  $\delta_H$  3.84 (6H, s)) and one hydroxyl group, combined with the observed optical rotation value of  $[\alpha]_D^{25} + 23.9$  ( $c$  0.1, MeOH) suggested that the structure of compound **5** was identified to be (+)-syringaresinol [11].

Compound **6** was obtained as an amorphous white powder. The molecular formula was identified to be  $\text{C}_{20}\text{H}_{20}\text{O}_6$  by its HR-ESIMS data at  $m/z$  357.1330  $[\text{M}+\text{H}]^+$  (calcd. for  $\text{C}_{20}\text{H}_{21}\text{O}_6^+$ , 357.1333) and 339.1228  $[\text{M}-\text{H}_2\text{O}+\text{H}]^+$  (calcd. for  $\text{C}_{20}\text{H}_{19}\text{O}_5^+$ , 339.1233). The  $^1\text{H}$ -NMR spectrum of **6** showed two olefinic protons at  $\delta_H$  7.63 and 6.69 with the coupling constant  $J = 15.6$  Hz featured for a *trans* configuration double bond; two protons located in *meta* position at  $\delta_H$  7.29 (1H, brs) and 7.24 (1H, d, 1.8 Hz); three protons of an ABX system (6.95 - 6.78 ppm); two methine groups at  $\delta_H$  5.61 (1H, d, 6.6 Hz) and 3.57 (1H, q, 6.0 Hz); a methylene group at  $\delta_H$  3.85 (2H, m); and two methoxy groups at  $\delta_H$  3.92 (3H, s) and 3.82 (3H, s). The  $^{13}\text{C}$ -NMR spectrum of **6** exhibited 20 carbons, including a carbonyl group at  $\delta_C$  196.1; four hydroxyl olefinic carbons (153.0 - 146.0 ppm); a double bond at  $\delta_C$  156.1 and  $\delta_C$  127.1; the oxygenated methine and methylene groups at  $\delta_C$  90.1 and 64.6, respectively; and two methoxy groups at  $\delta_C$  56.8 and 56.4. In the HMBC spectrum, proton  $\delta_H$  3.57 (H-8) illustrated correlations with  $\delta_C$  153.0 (C-4') and 120.0 (C-6'), proving that two  $\text{C}_6\text{-C}_3$  units were connected through C-8 and C-5'. In addition, proton  $\delta_H$  5.61 (H-7) interacted with  $\delta_C$  153.0 (C-4') and 131.3 (C-5'), suggesting the formation of a furan ring. The optical rotation of compound **6** was measured to give  $[\alpha]_D^{25} + 10.0$  ( $c$  0.1, MeOH). Detailed spectral data analyses and comparison with the published reference [12] led to the structure elucidation as balanophonin.

To the best of our knowledge, these six compounds were isolated and structure elucidated for the first time from the genus *Cansjera*. Two lignan esters, alyterinate A and B, were reported from *Alyxia schlechteri* and *Willughbeia coriacea*. Alyterinate A showed cytotoxic activity against HeLa cell line [13]. (+)-Pinoresinol, (+)-medioresinol, and (+)-syringaresinol were commonly found in foods and medicinal plants, which exhibited anti-inflammatory effects by influencing the NF- $\kappa$ B and MAPK pathways, demonstrated antitumor activities through the regulation of apoptosis, and alleviated the progression of osteoporosis [14]. Balanophonin, a dihydrobenzofuran neolignan, was investigated to

illustrate the anti-neuroinflammatory [15], anti-HIV-1 [16], and PTP1B inhibitory [12] activities. This study provides further information about the

phytochemical composition of *C. rheedei*, which contributes to the scientific understanding of this herb in usage and quality control.



**Fig.2.** Key HMBC and COSY correlations of the isolated compounds (1–6)

#### 4. Conclusion

Five lignans and one neolignan were isolated from the stems of *C. rheedei* J.F.Gmel.. The structures of these compounds were elucidated by spectral data analyses and by comparison with

those in references as alyterinate A (1), alyterinate B (2), (+)-pinioresinol (3), (+)-medioresinol (4), (+)-syringaresinol (5), and balanophonin (6). All these chemical constituents were reported for the first time from the genus *Cansjera*.

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