

*Supplementary Material*

# Design, Synthesis of Novel Tetrandrine-14-L-Amino Acid and Tetrandrine-14-L-Amino Acid-Urea Derivatives as Potential Anti-cancer Agents

Sheng-Cao Hu <sup>1,2,3,†</sup>, Jin Yang <sup>1,3,†</sup>, Chao Chen <sup>2,3</sup>, Jun-Rong Song <sup>2,3,\*</sup> and Wei-Dong Pan <sup>1,2,3,\*</sup>

<sup>1</sup> College of Pharmacy, Zunyi Medical University, Zunyi 563000, PR China; hushengcao0221@163.com (S.-C.H.); jinyangtrcwsys@sina.com (J.Y.)

<sup>2</sup> State Key Laboratory of Functions and Applications of Medicinal Plants, Guizhou Medical University, Guiyang 550014, PR China; cc283818640@163.com

<sup>3</sup> The Key Laboratory of Chemistry for Natural Products of Guizhou Province and Chinese Academy of Sciences, Guiyang 550014, PR China

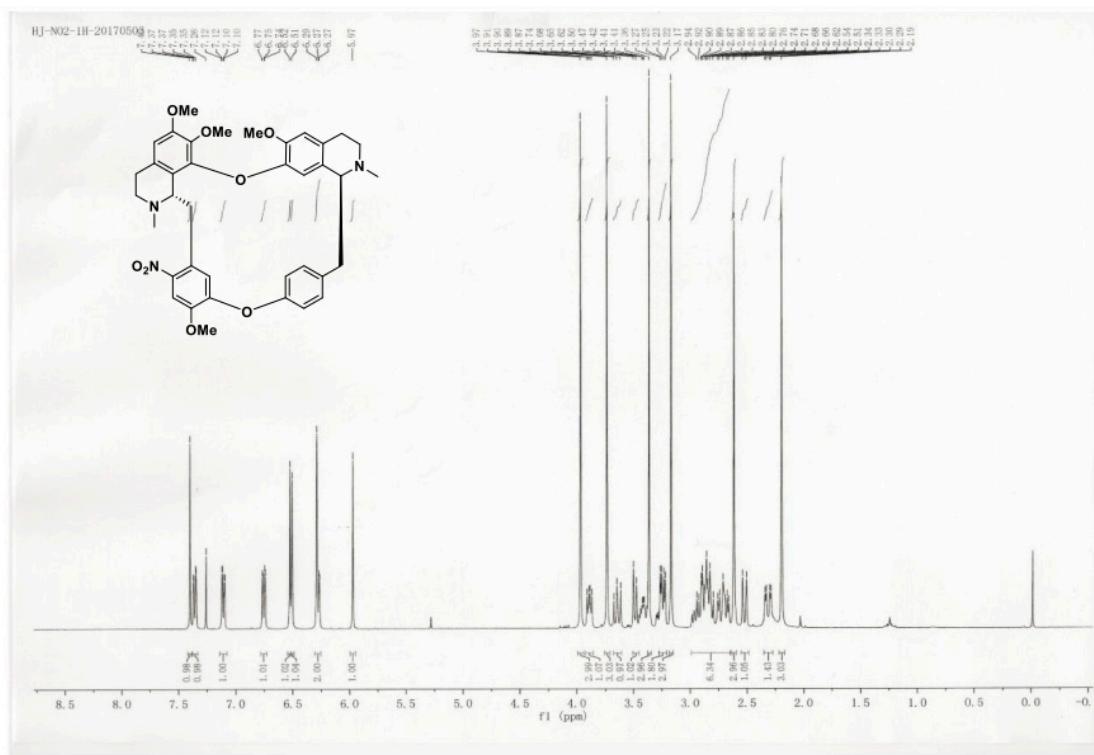
\* Correspondence: wdpn@163.com (W.-D.P.), 18275365116@163.com (J.-R.S.), Tel: +86 18985130307 (W.-D.P.)

† These authors contributed equally to this work.

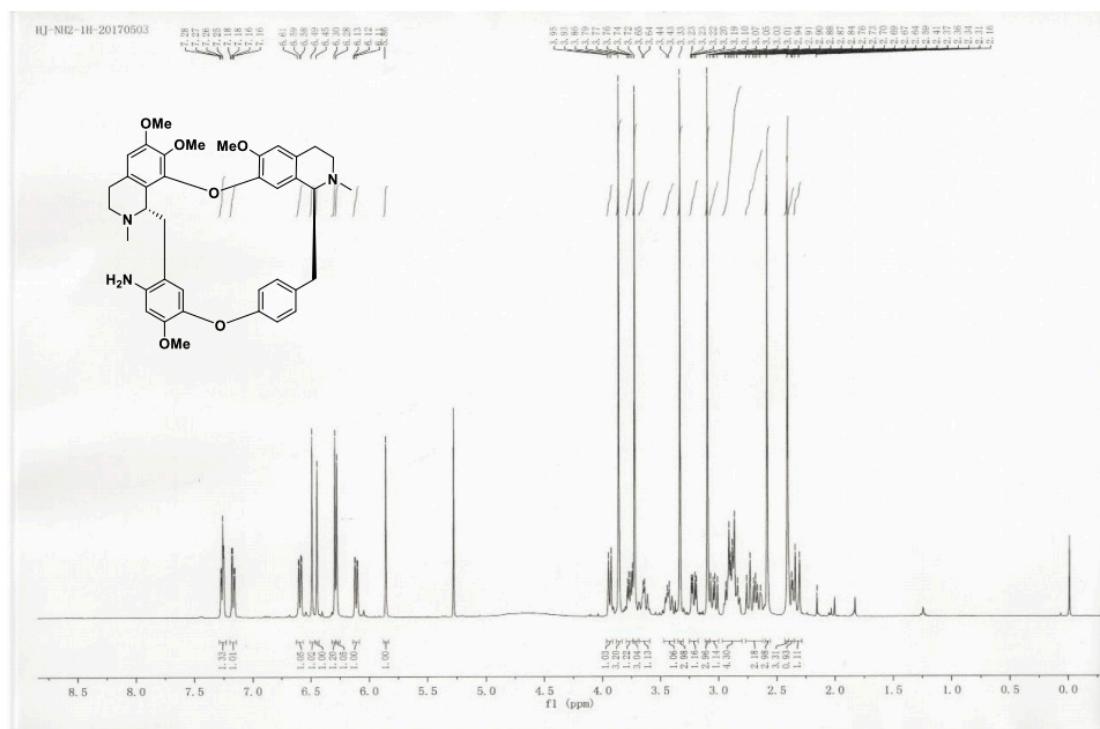
## Table of contents

1. <sup>1</sup> H-NMR Spectra of Tet-NO <sub>2</sub> , Tet-NH <sub>2</sub> and <b>1a-3k</b> .....	2
2. <sup>13</sup> C-NMR Spectra of Tet-NO <sub>2</sub> , Tet-NH <sub>2</sub> and <b>1a-3k</b> .....	15

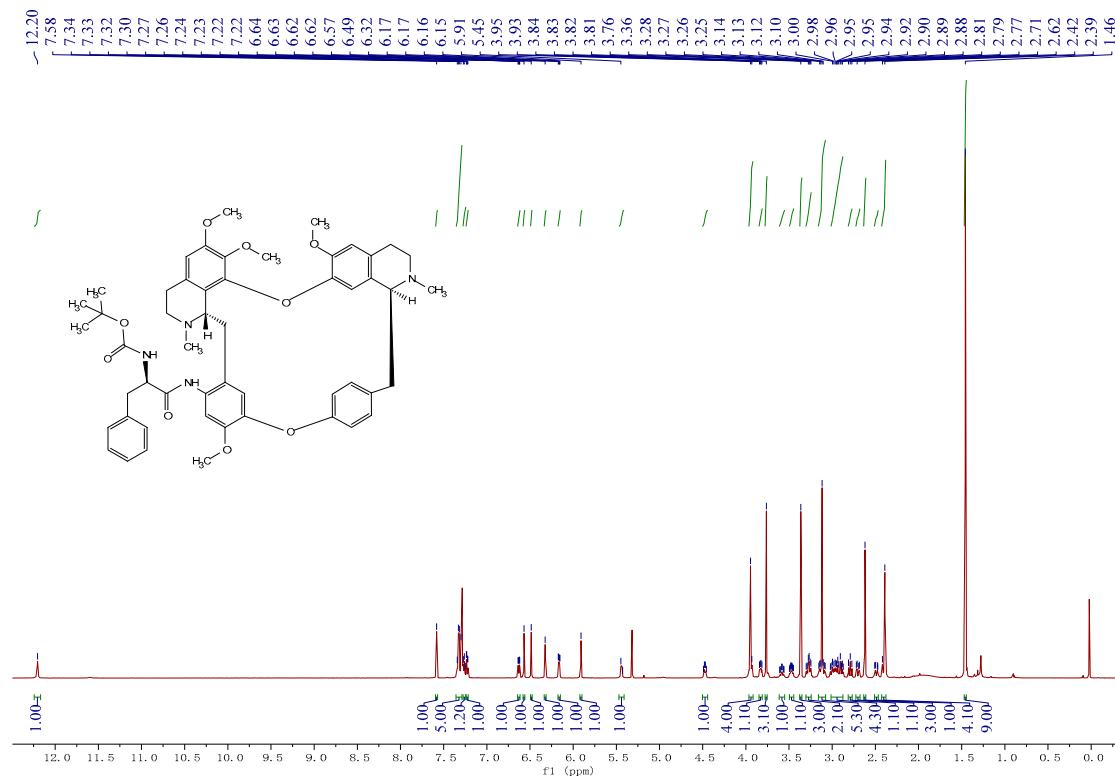
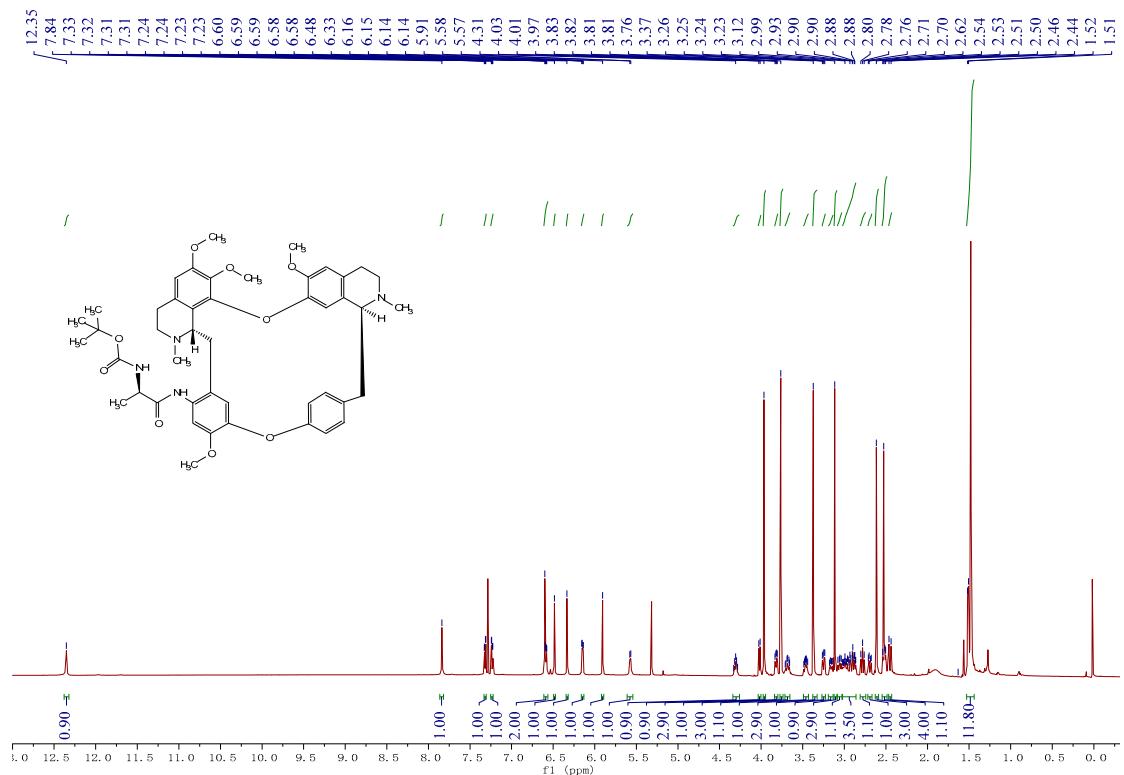
## 1. $^1\text{H}$ -NMR Spectra of 1a-3k

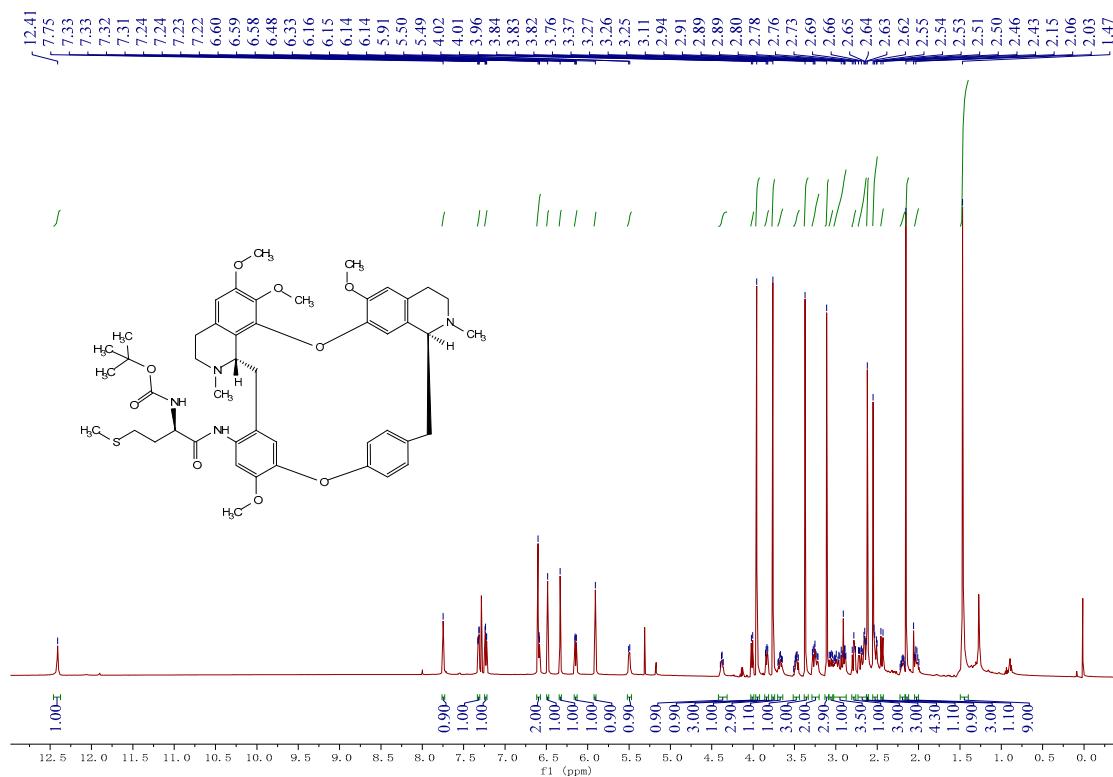


**Figure S1.**  $^1\text{H}$ -NMR Spectra of Tet-NO<sub>2</sub> in CDCl<sub>3</sub>.

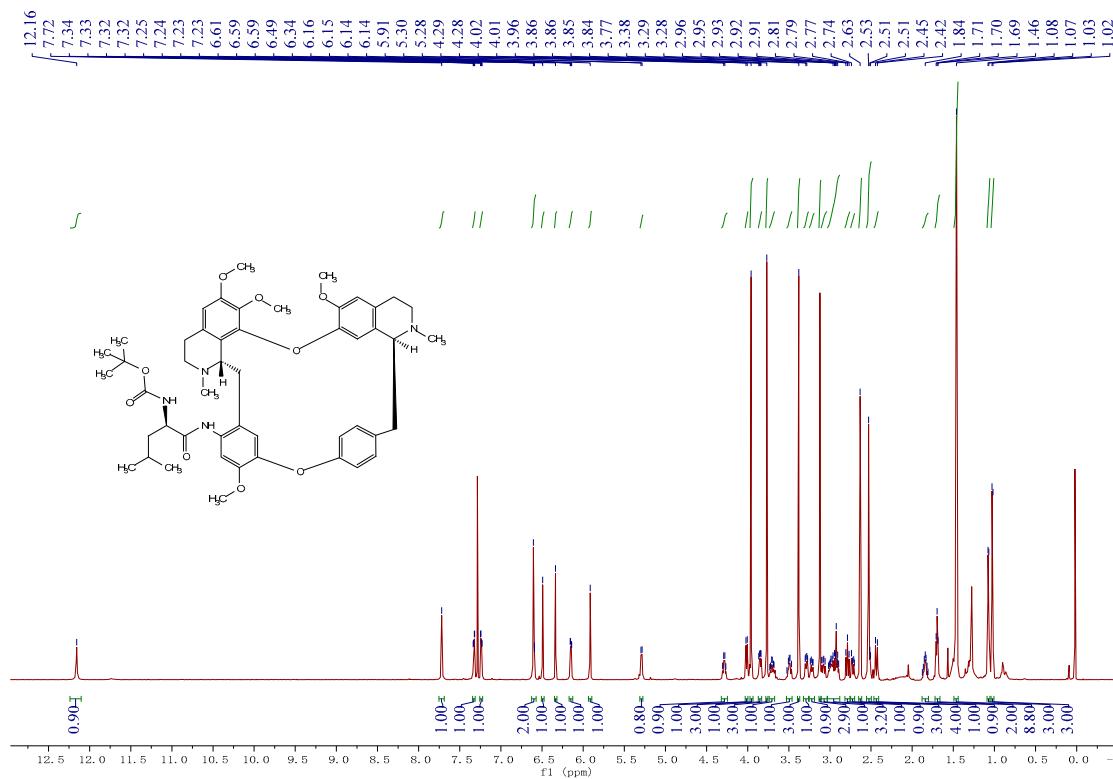


**Figure S2.**  $^1\text{H}$ -NMR Spectra of Tet-NH<sub>2</sub> in CDCl<sub>3</sub>.

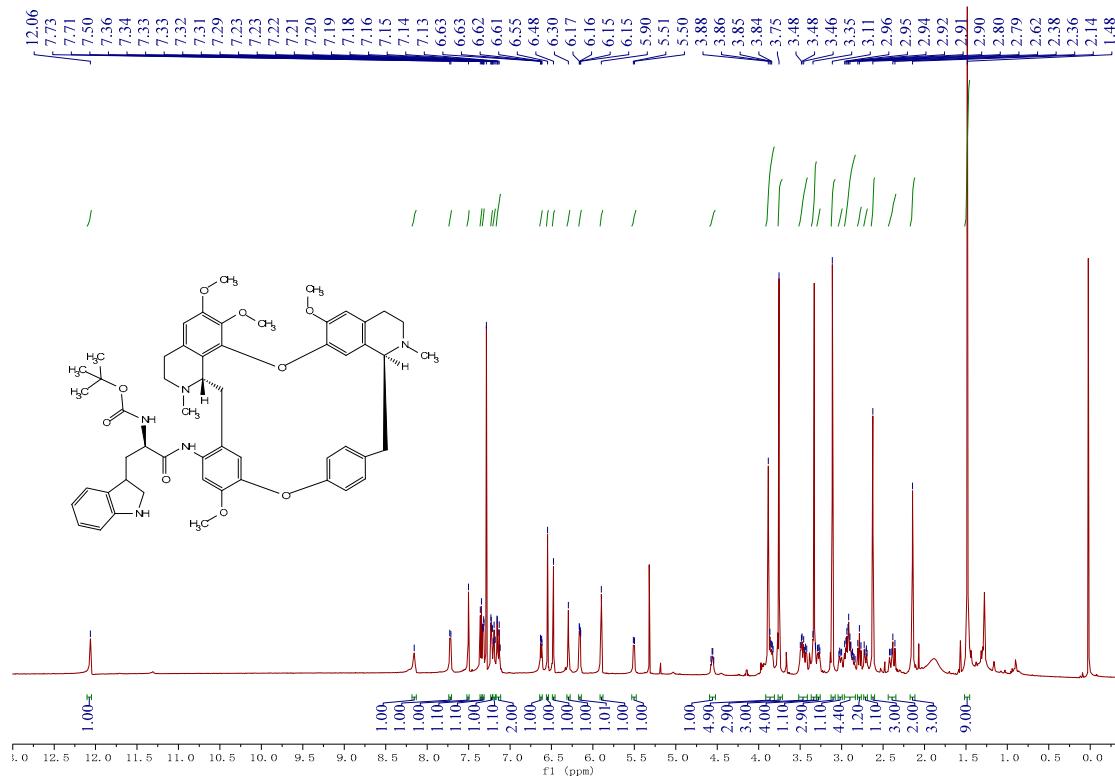
**Figure S3.**  $^1\text{H}$ -NMR Spectra of **1a** in  $\text{CDCl}_3$ .**Figure S4.**  $^1\text{H}$ -NMR Spectra of **1b** in  $\text{CDCl}_3$ .



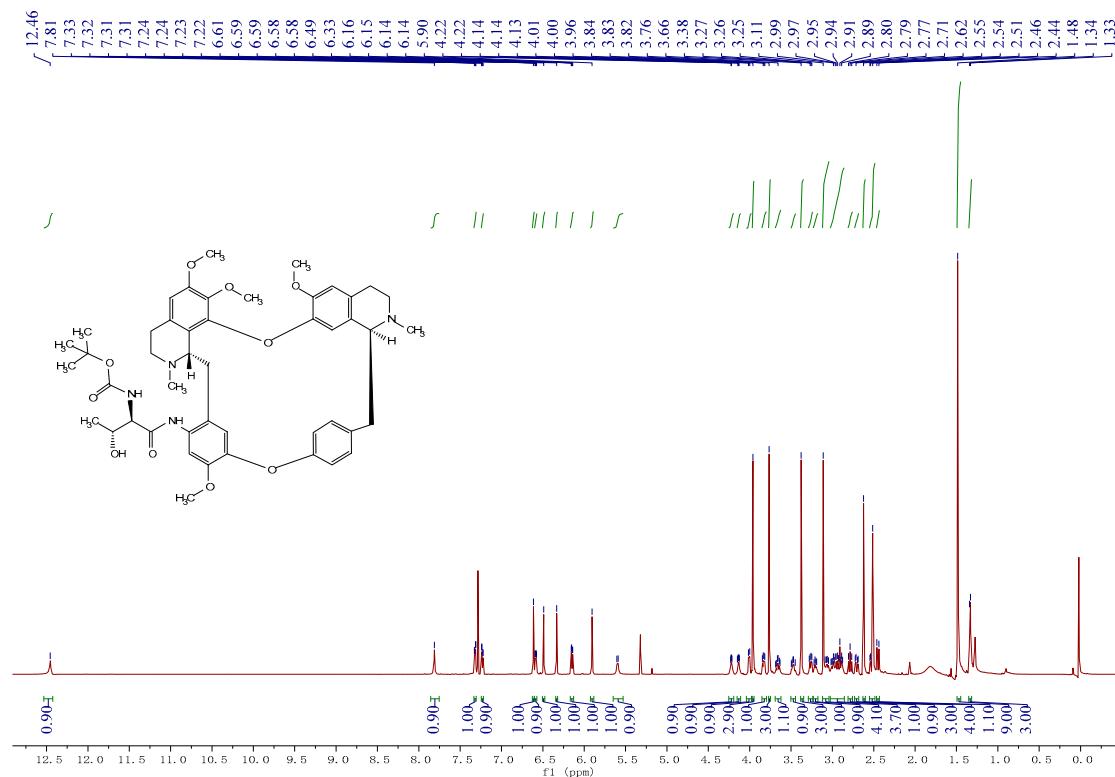
**Figure S5.**  $^1\text{H}$ -NMR Spectra of **1c** in  $\text{CDCl}_3$ .



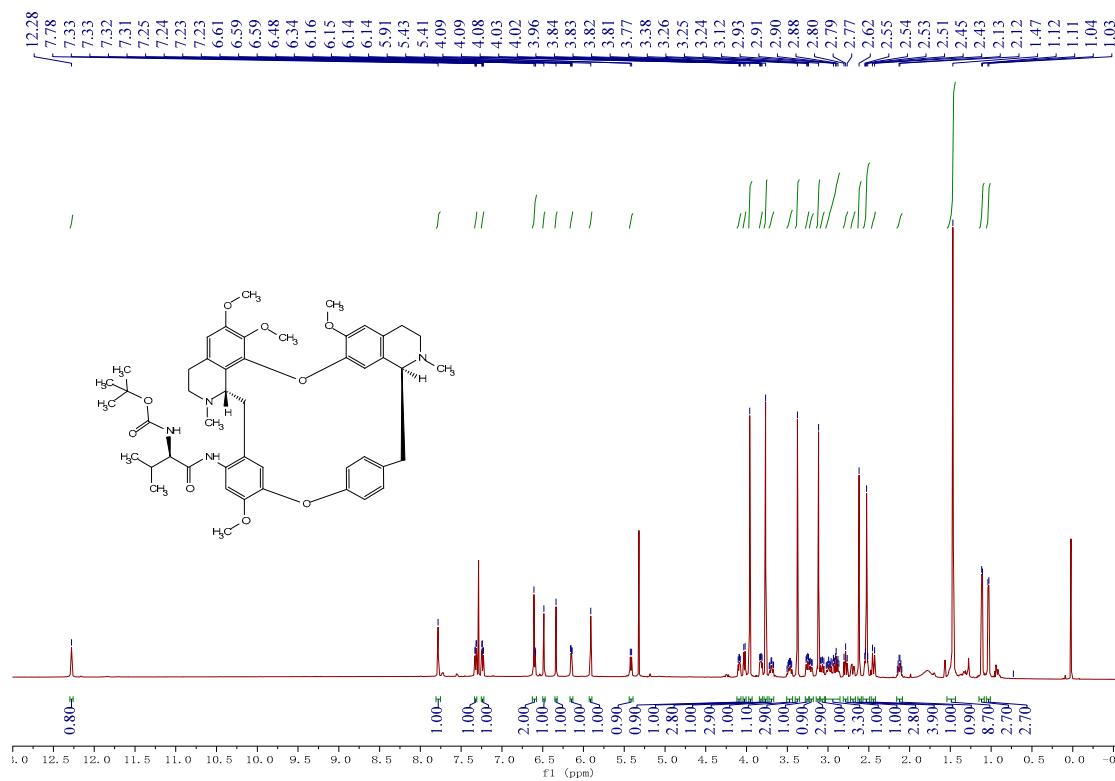
**Figure S6.**  $^1\text{H}$ -NMR Spectra of **1d** in  $\text{CDCl}_3$ .



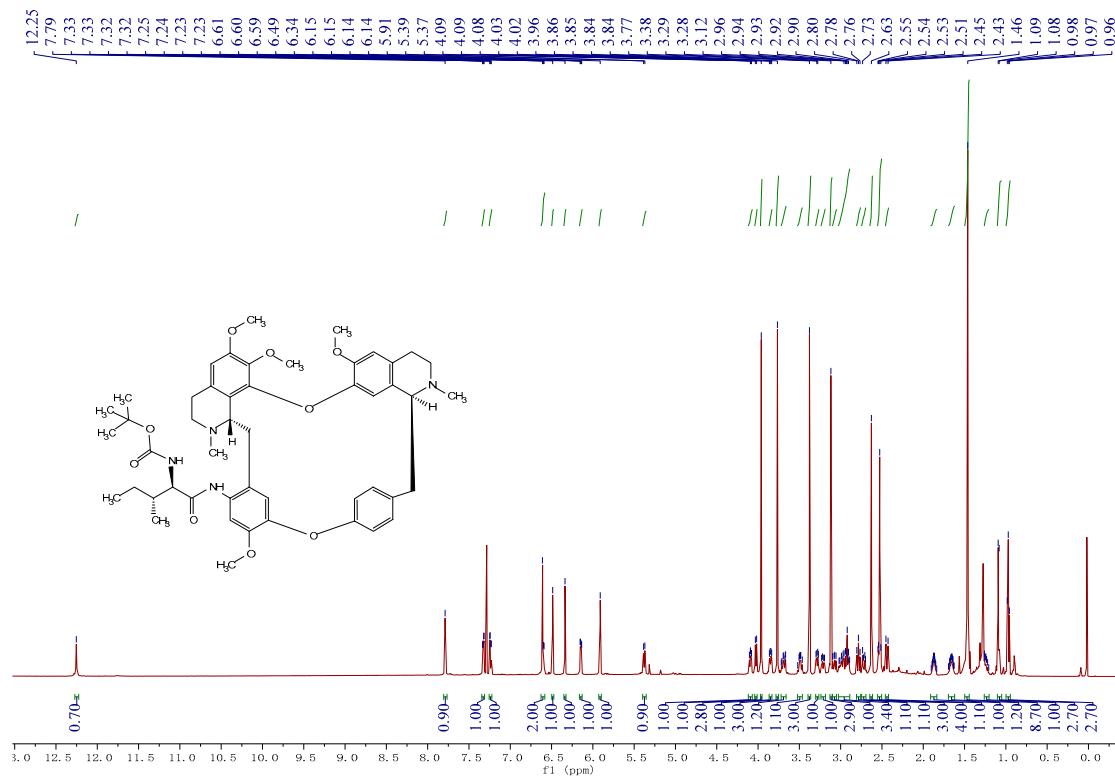
**Figure S7.**  $^1\text{H}$ -NMR Spectra of **1e** in  $\text{CDCl}_3$ .



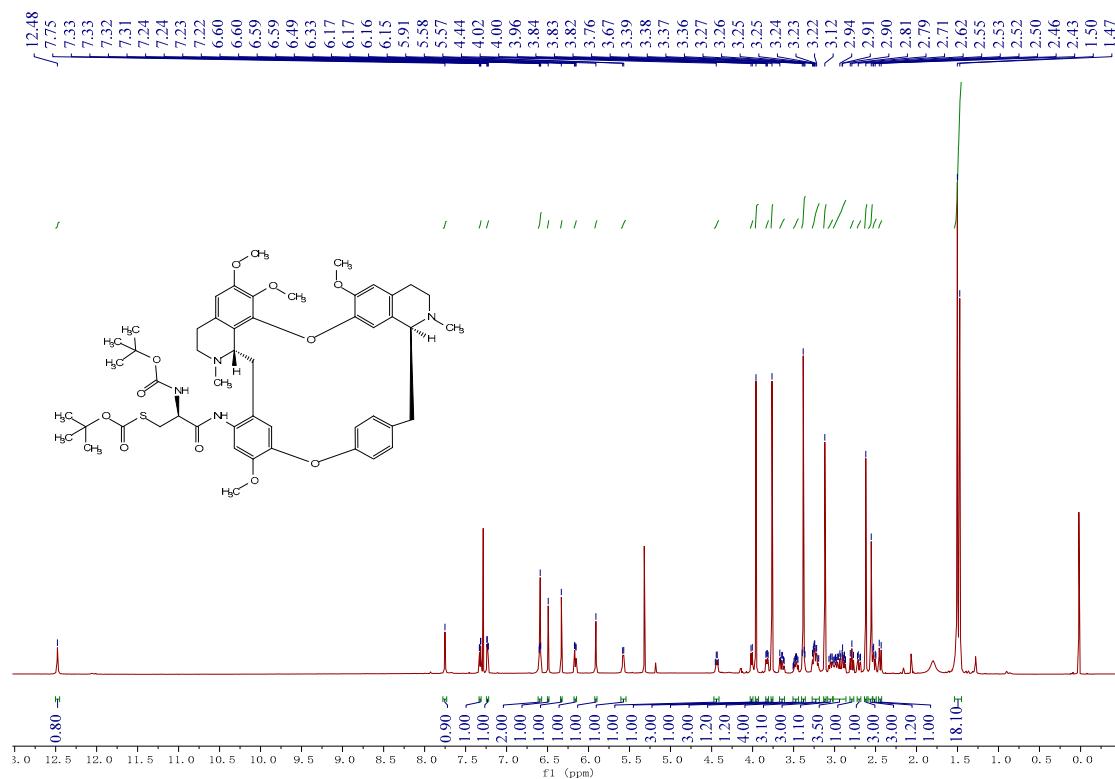
**Figure S8.**  $^1\text{H}$ -NMR Spectra of **1f** in  $\text{CDCl}_3$ .



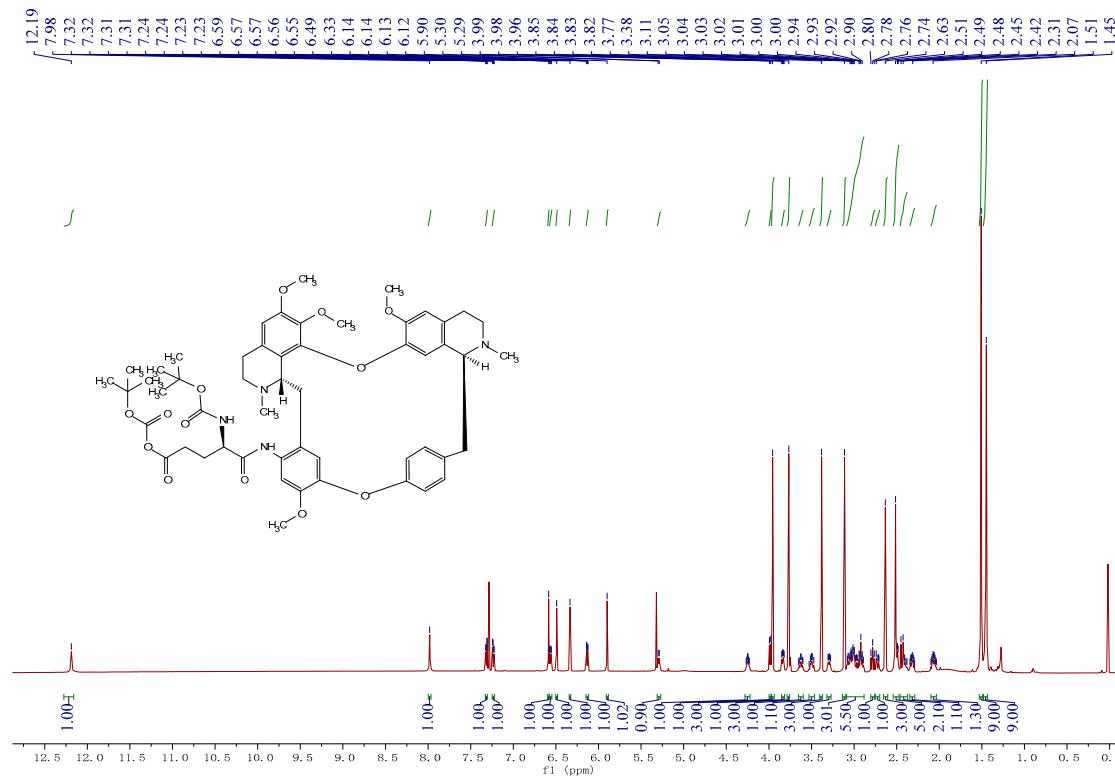
**Figure S9.**  $^1\text{H}$ -NMR Spectra of **1g** in  $\text{CDCl}_3$ .



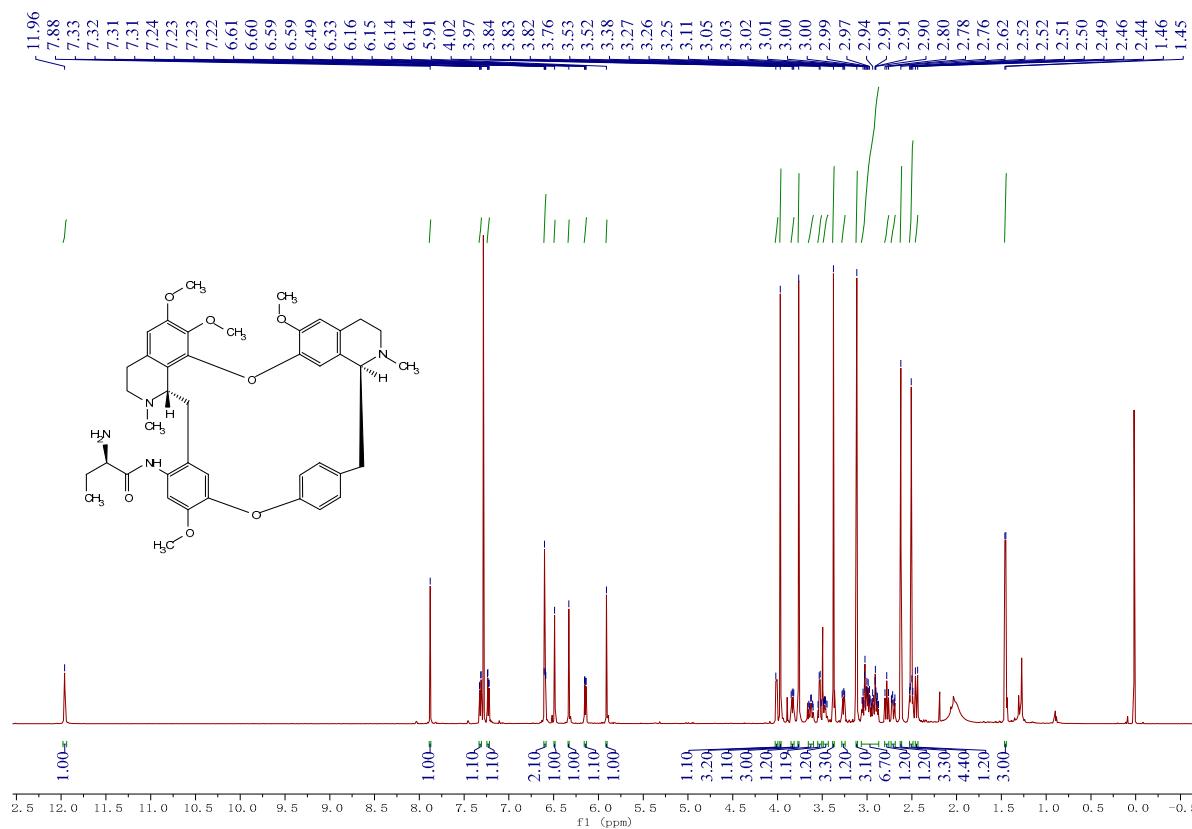
**Figure S10.**  $^1\text{H}$ -NMR Spectra of **1h** in  $\text{CDCl}_3$ .



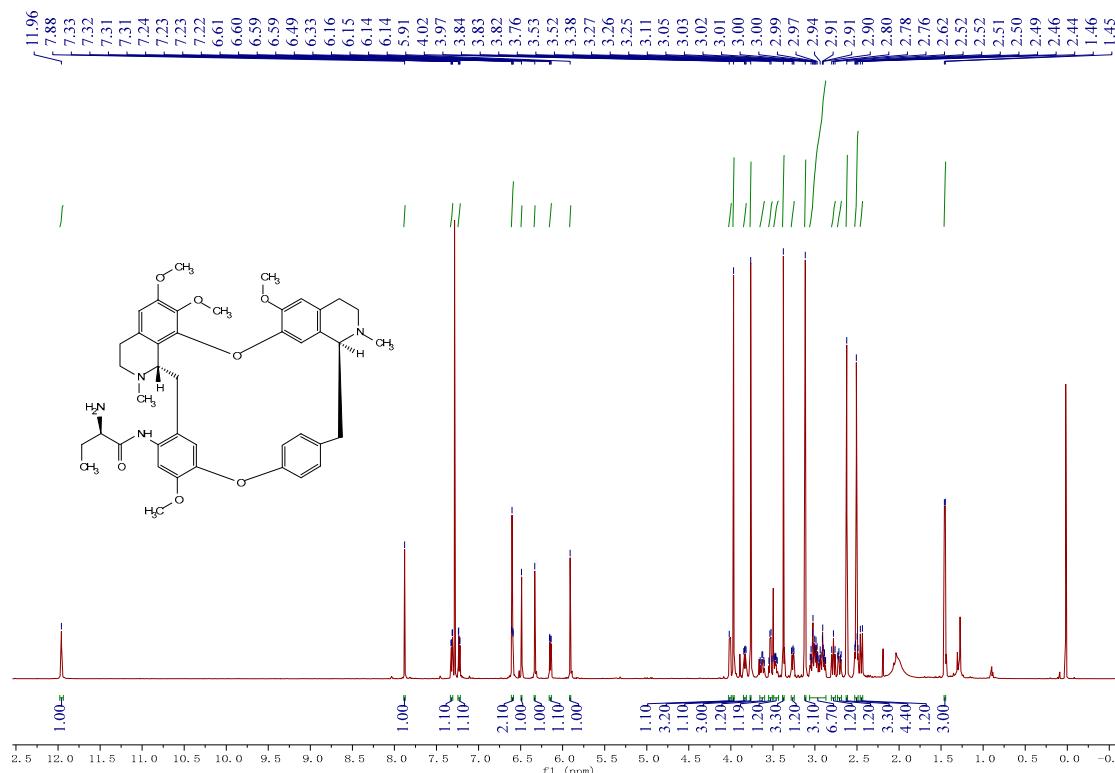
**Figure S11.**  $^1\text{H}$ -NMR Spectra of **1i** in  $\text{CDCl}_3$ .



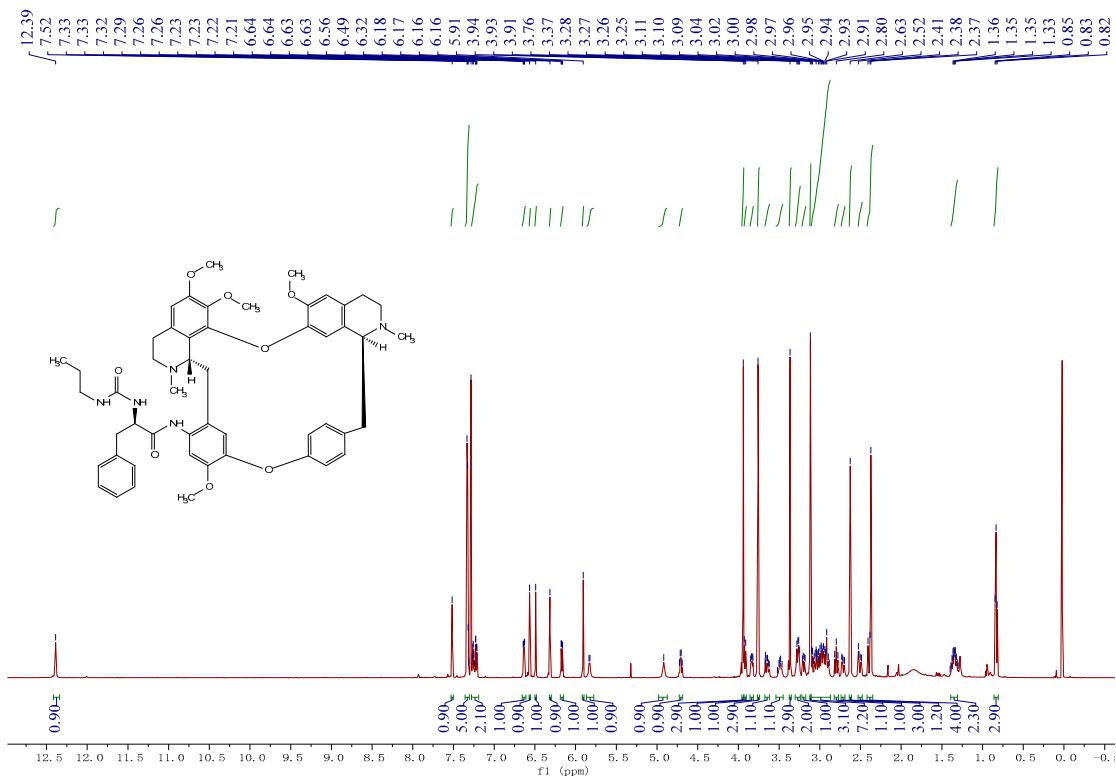
**Figure S12.**  $^1\text{H}$ -NMR Spectra of **1j** in  $\text{CDCl}_3$ .



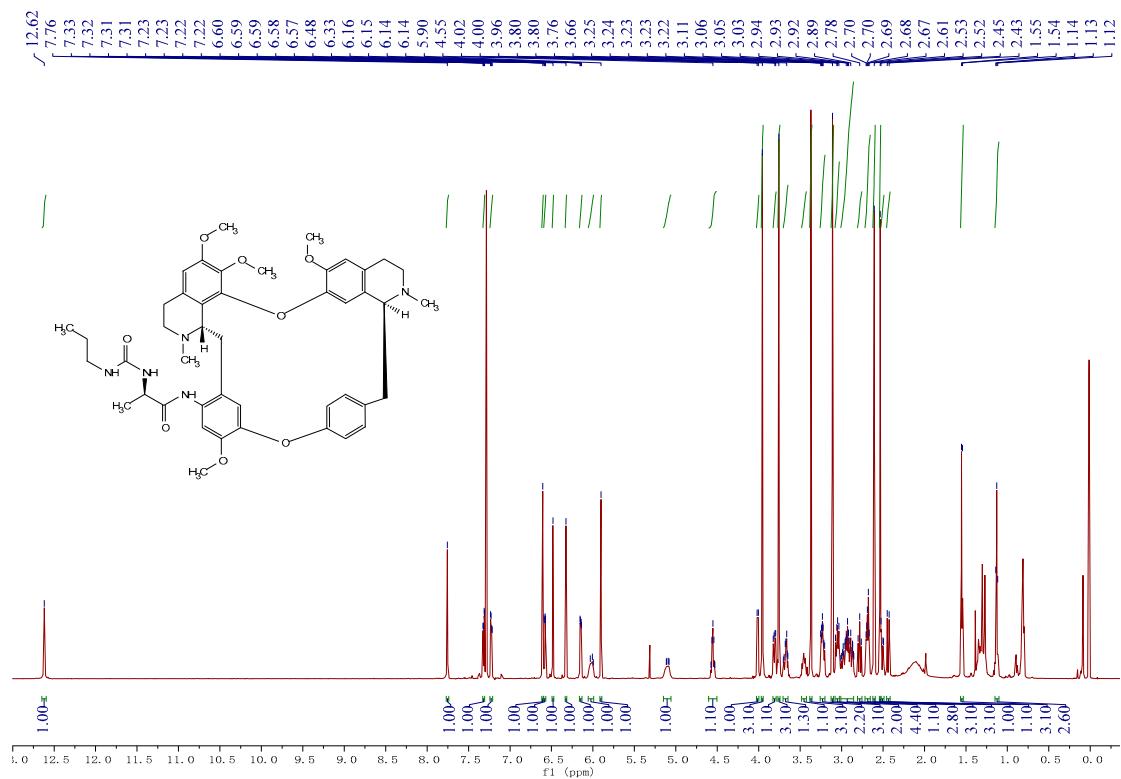
**Figure S13.**  $^1\text{H}$ -NMR Spectra of **1k** in  $\text{CDCl}_3$ .



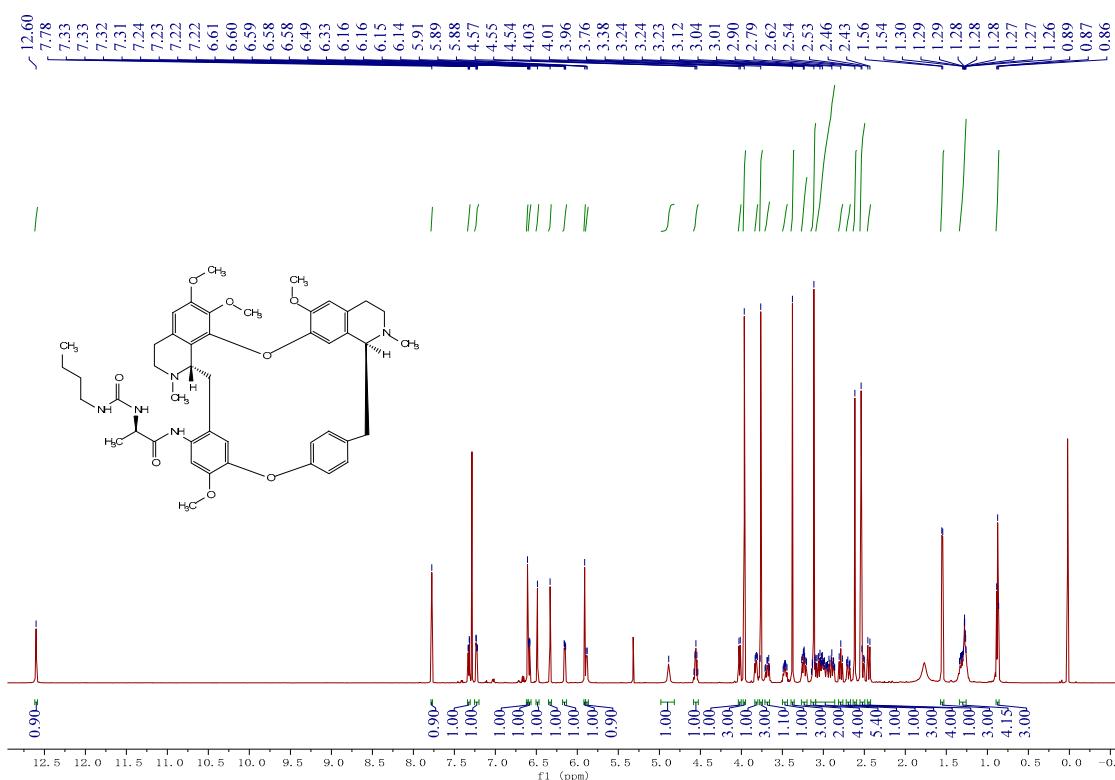
**Figure S14.**  $^1\text{H}$ -NMR Spectra of **1I** in  $\text{CDCl}_3$ .



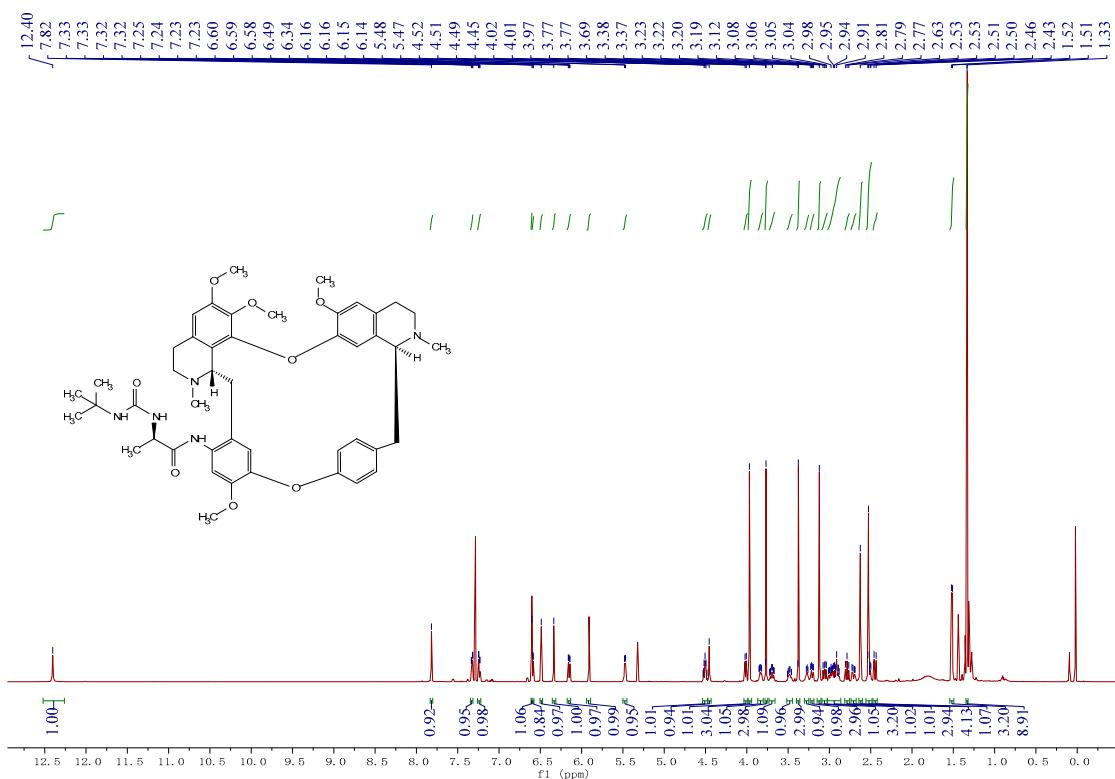
**Figure S15.**  $^1\text{H}$ -NMR Spectra of **2a** in  $\text{CDCl}_3$ .



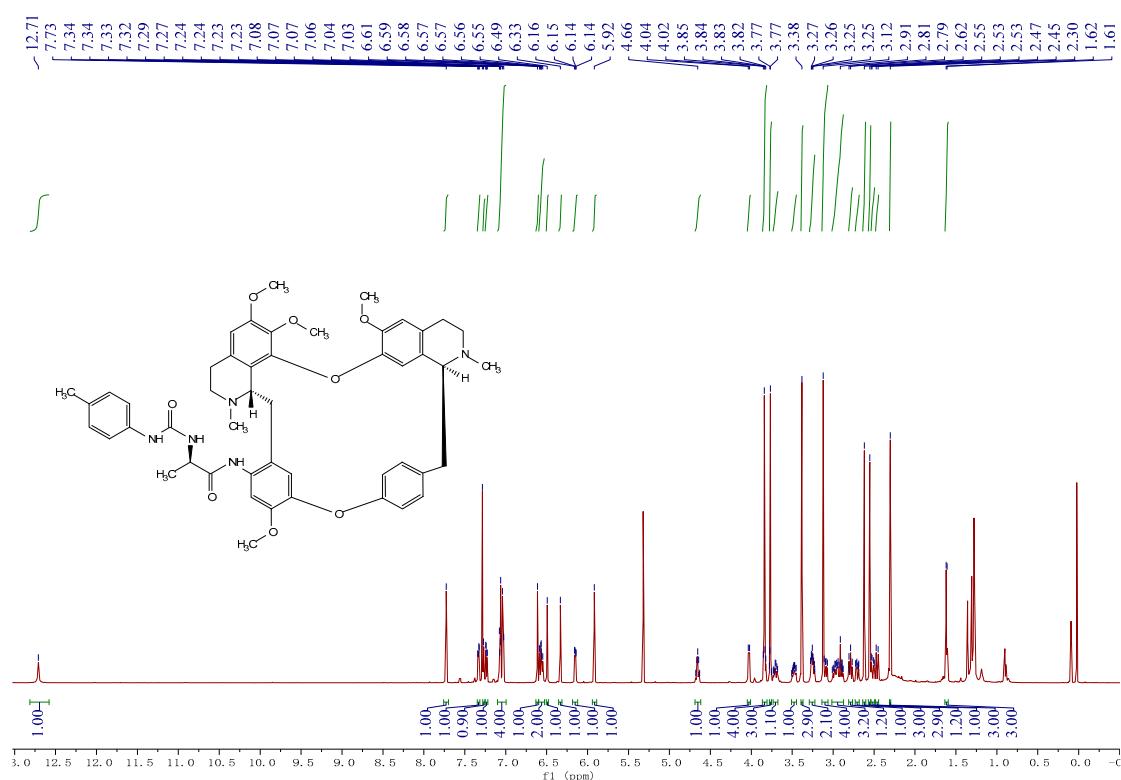
**Figure S16.**  $^1\text{H}$ -NMR Spectra of **3a** in  $\text{CDCl}_3$ .



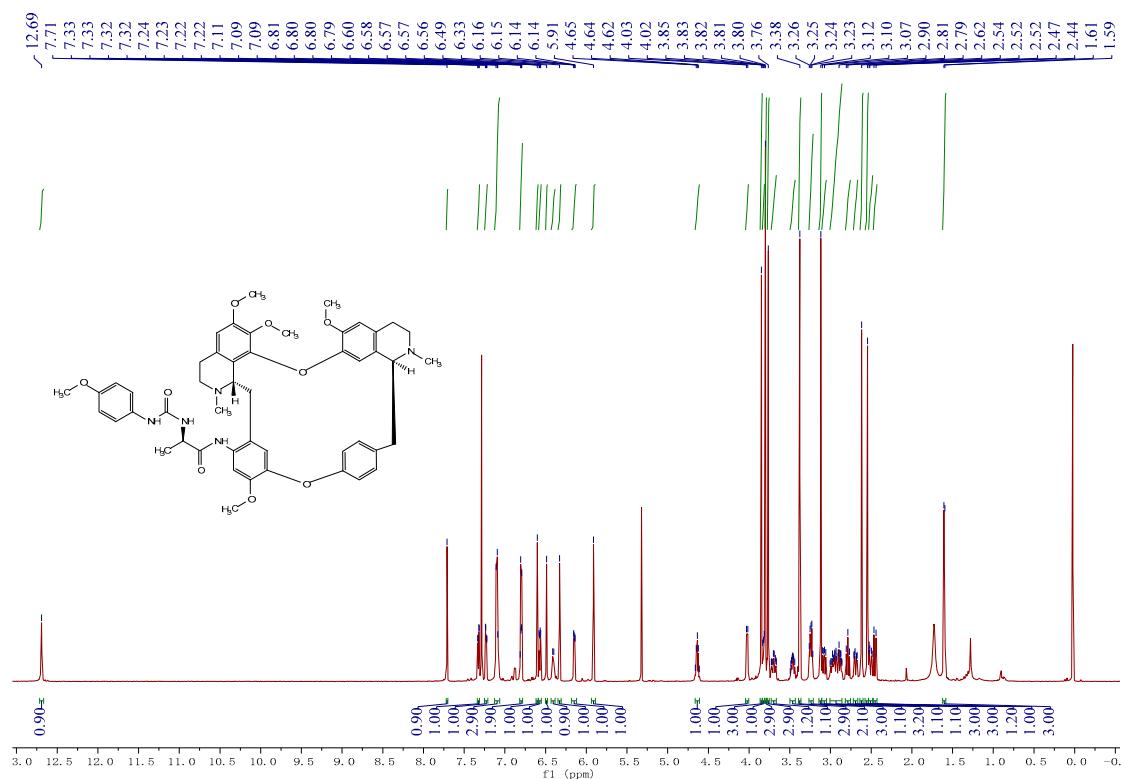
**Figure S17.**  $^1\text{H}$ -NMR Spectra of **3b** in  $\text{CDCl}_3$ .



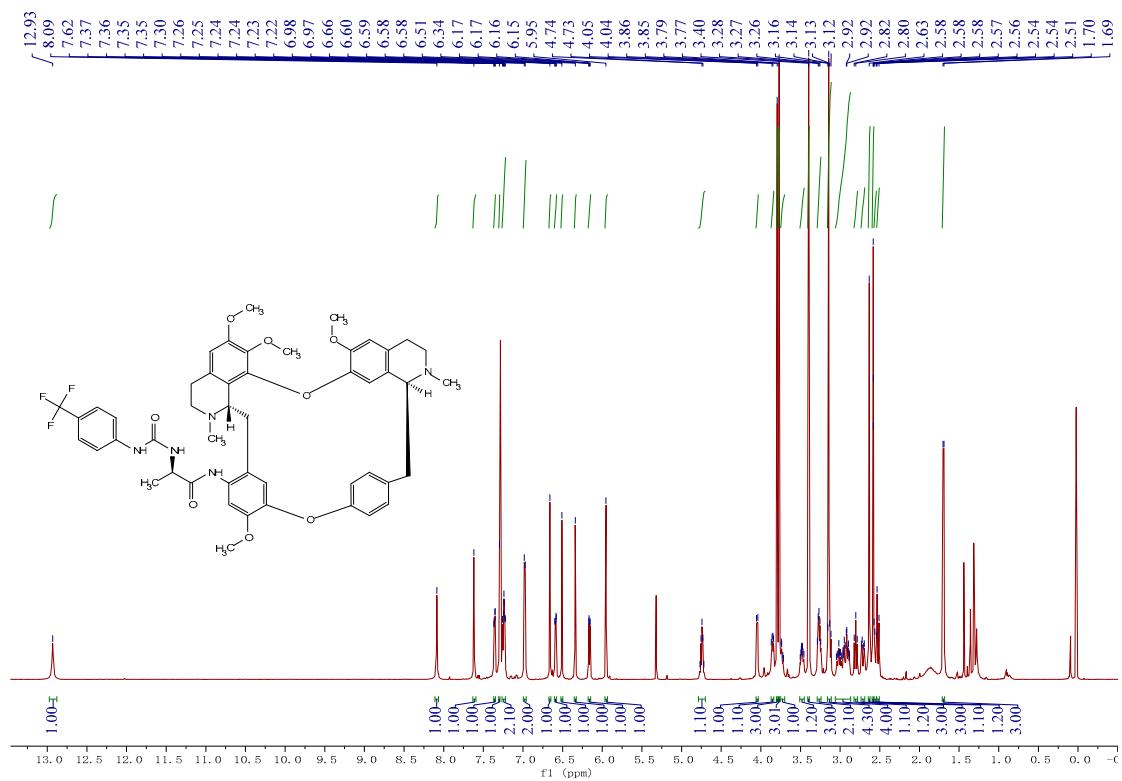
**Figure S18.**  $^1\text{H}$ -NMR Spectra of **3c** in  $\text{CDCl}_3$ .



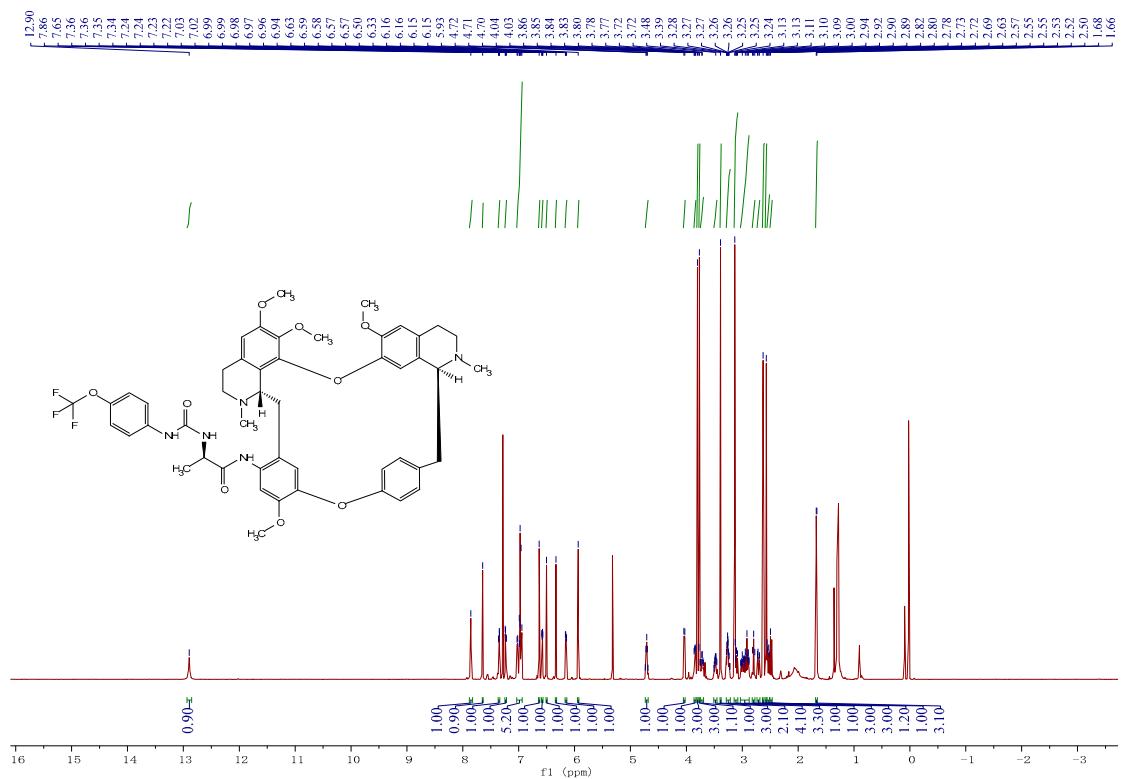
**Figure S19.**  $^1\text{H}$ -NMR Spectra of **3d** in  $\text{CDCl}_3$ .



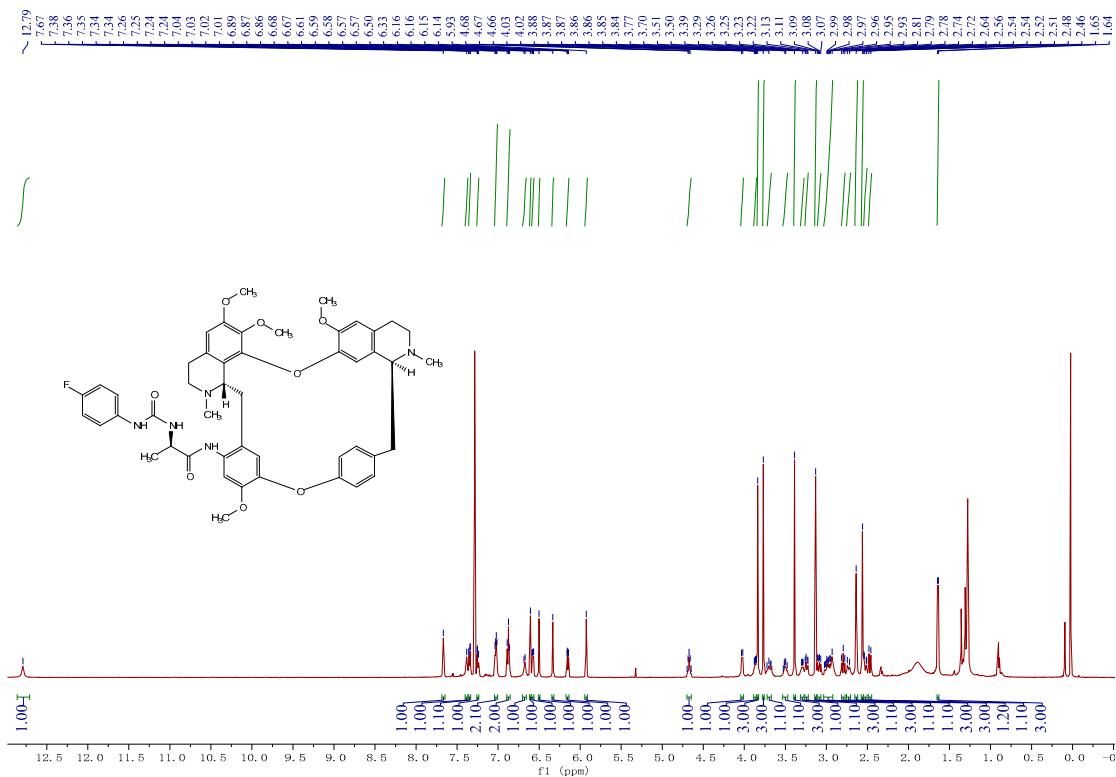
**Figure S20.**  $^1\text{H}$ -NMR Spectra of **3e** in  $\text{CDCl}_3$ .



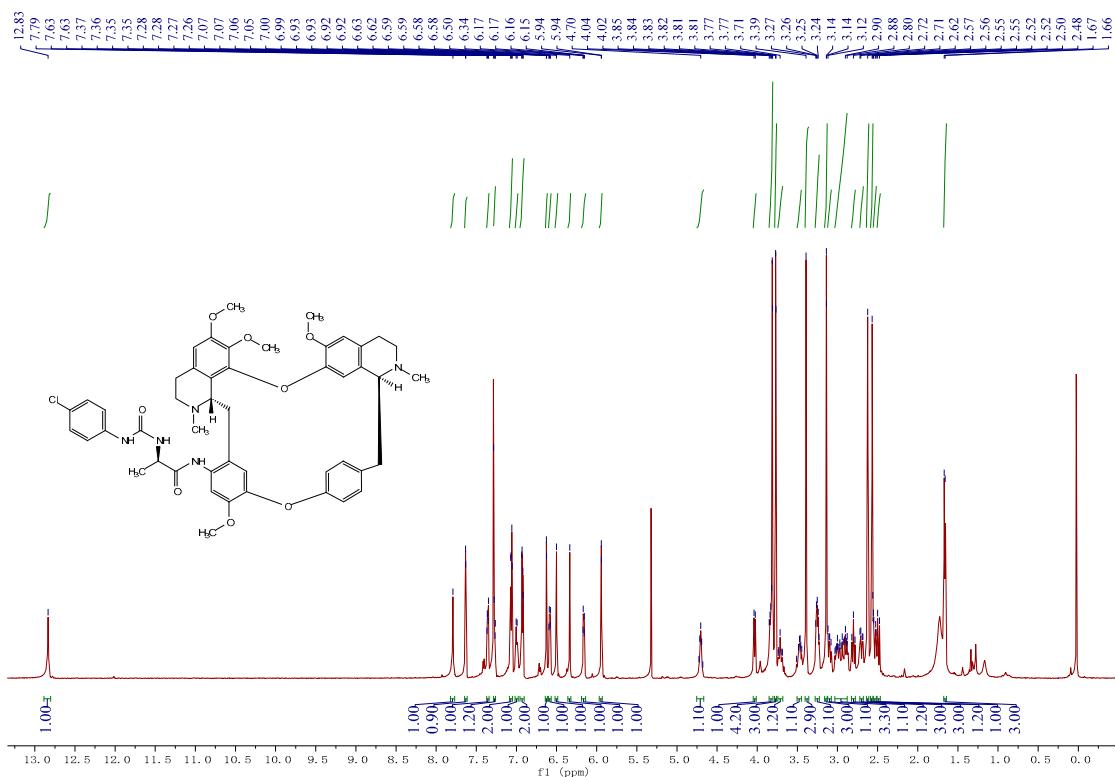
**Figure S21.**  $^1\text{H}$ -NMR Spectra of **3f** in  $\text{CDCl}_3$ .



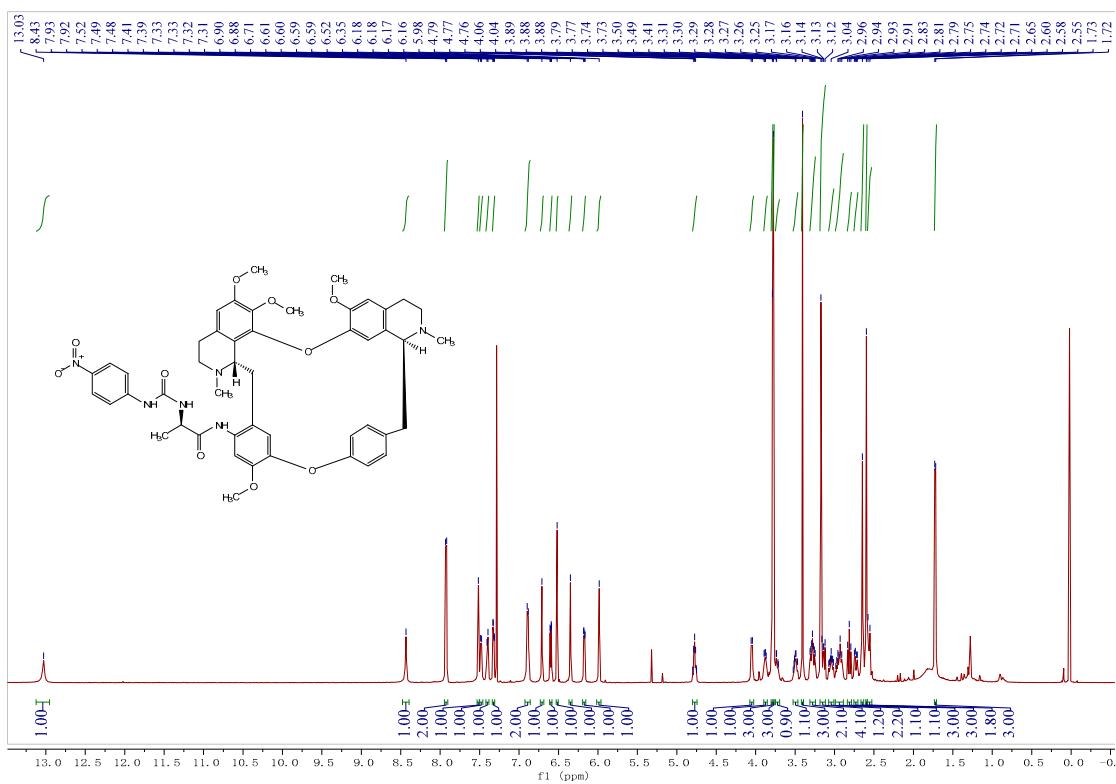
**Figure S22.**  $^1\text{H}$ -NMR Spectra of **3g** in  $\text{CDCl}_3$ .



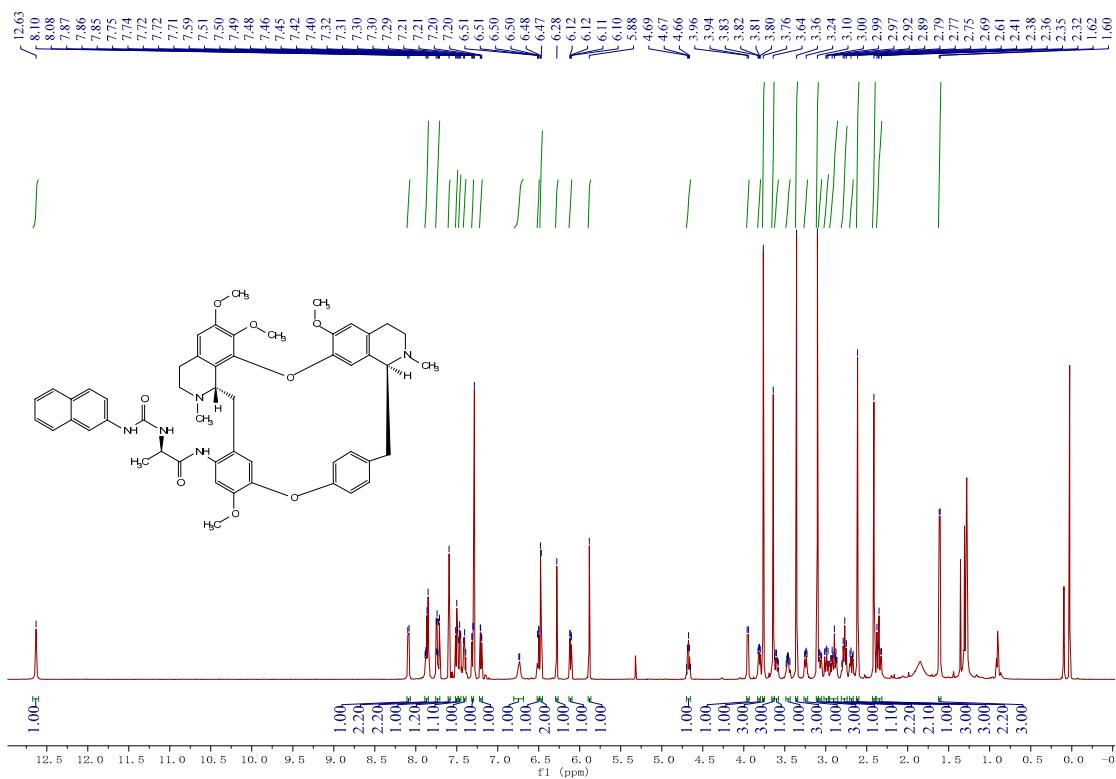
**Figure S23.**  $^1\text{H}$ -NMR Spectra of **3h** in  $\text{CDCl}_3$ .



**Figure S24.**  $^1\text{H}$ -NMR Spectra of **3i** in  $\text{CDCl}_3$ .

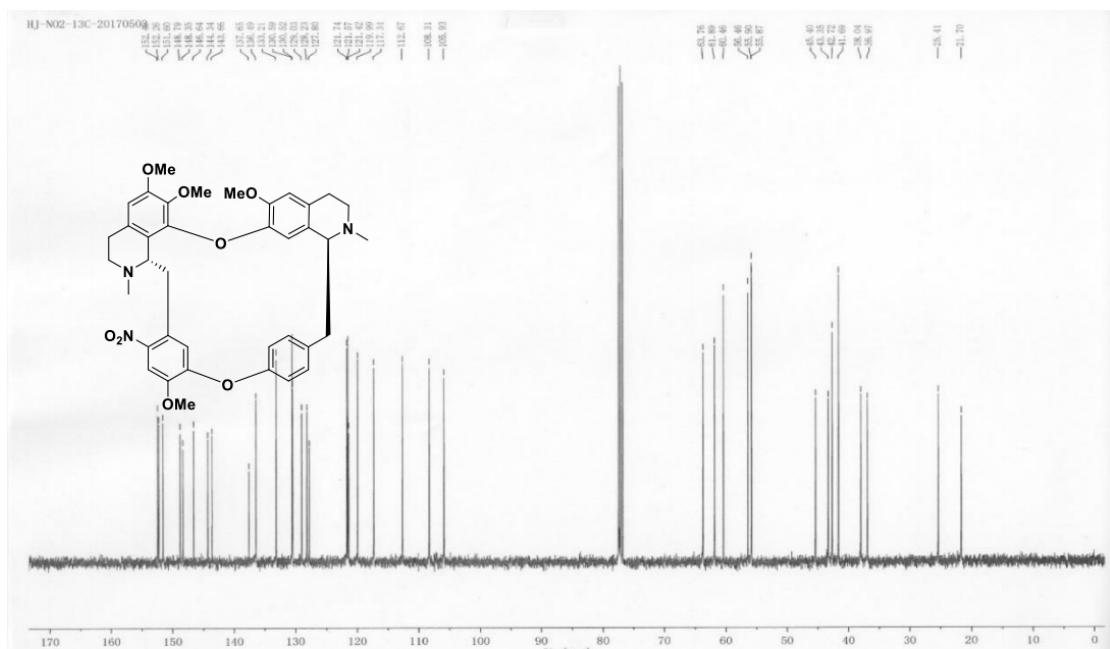


**Figure S25.**  $^1\text{H}$ -NMR Spectra of **3j** in  $\text{CDCl}_3$ .

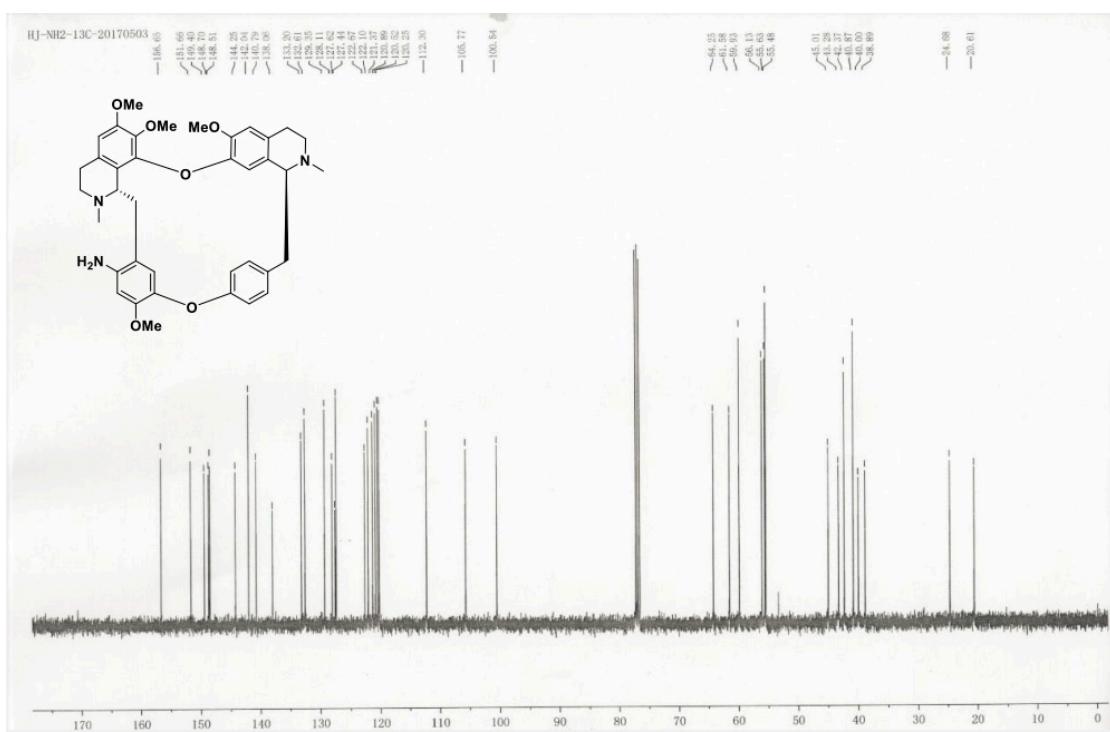


**Figure S26.**  $^1\text{H}$ -NMR Spectra of **3k** in  $\text{CDCl}_3$ .

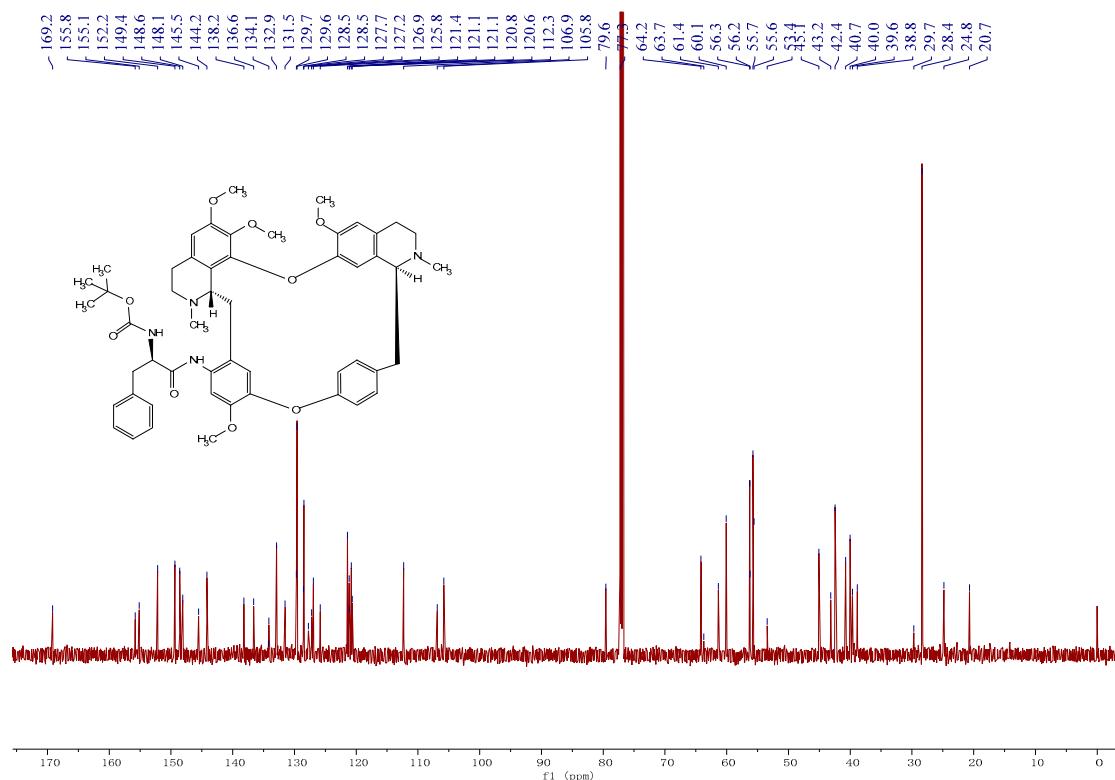
## 2. $^{13}\text{C}$ -NMR Spectra of 1a-3k



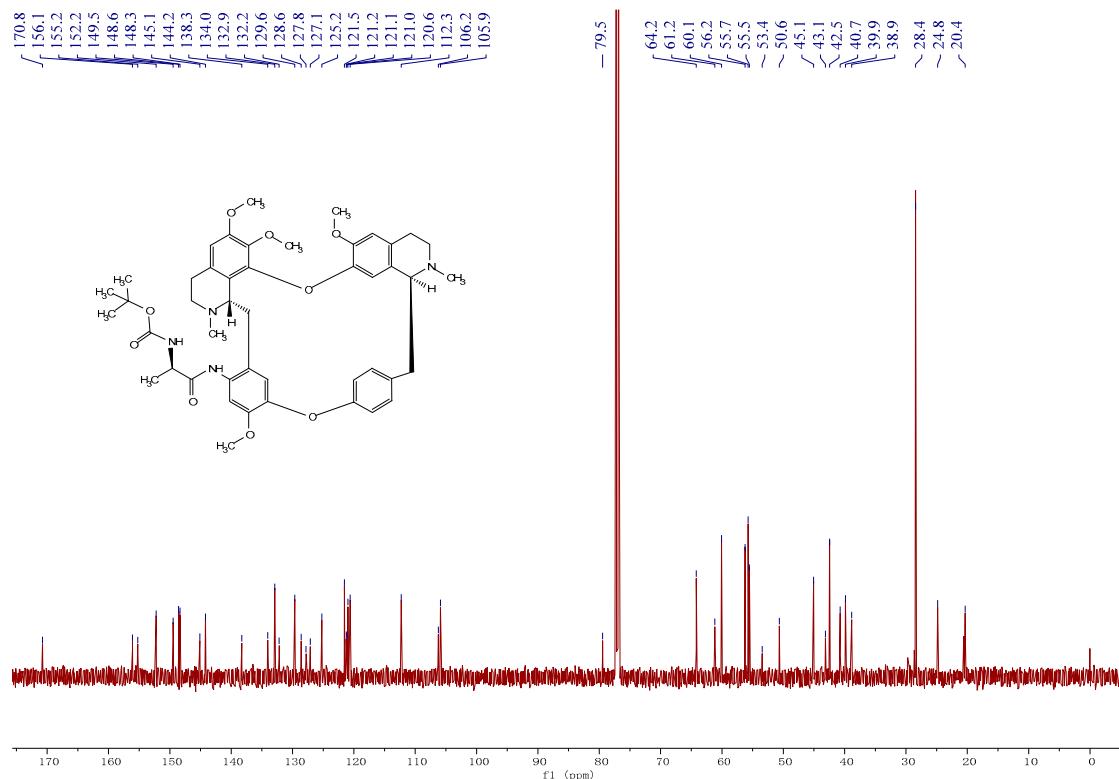
**Figure S27.**  $^{13}\text{C}$ -NMR Spectra of Tet-NO<sub>2</sub> in CDCl<sub>3</sub>.



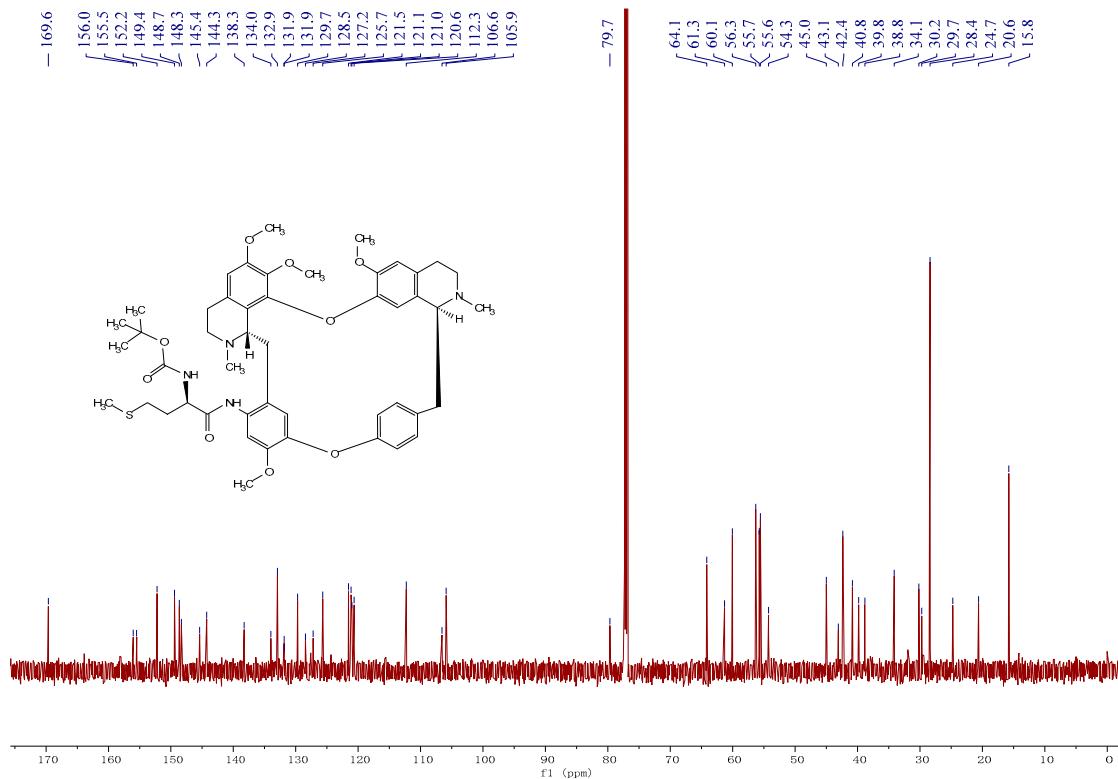
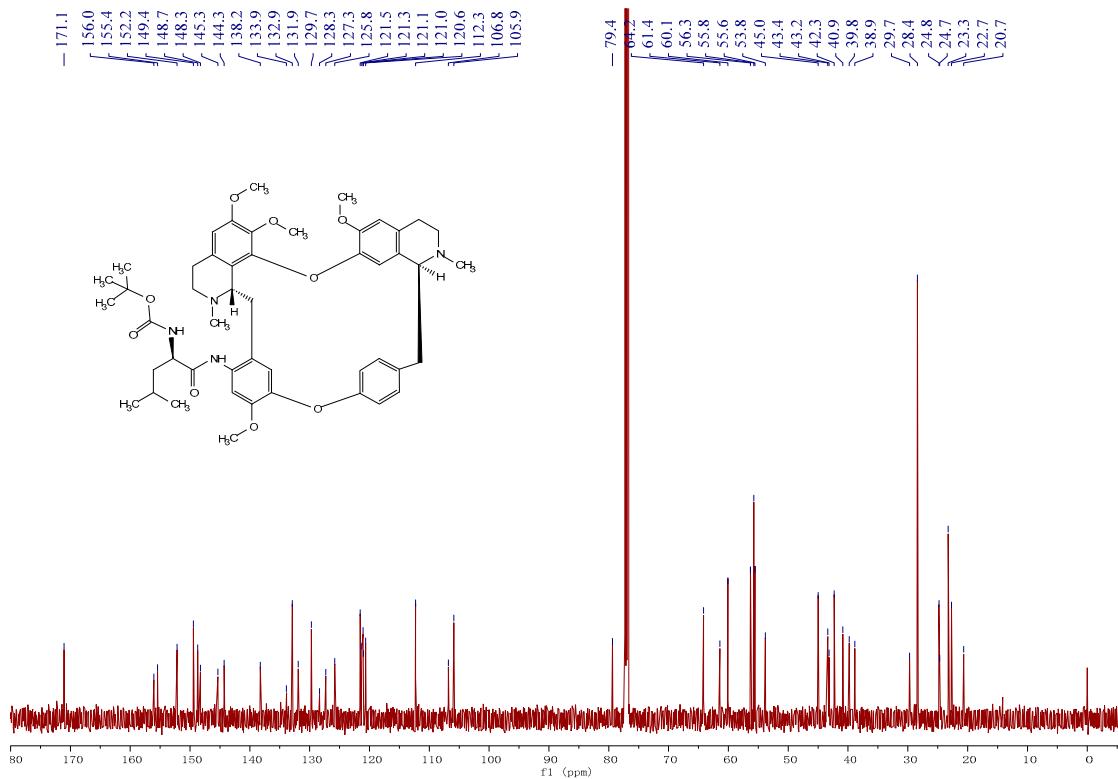
**Figure S28.**  $^{13}\text{C}$ -NMR Spectra of Tet-NH<sub>2</sub> in CDCl<sub>3</sub>.

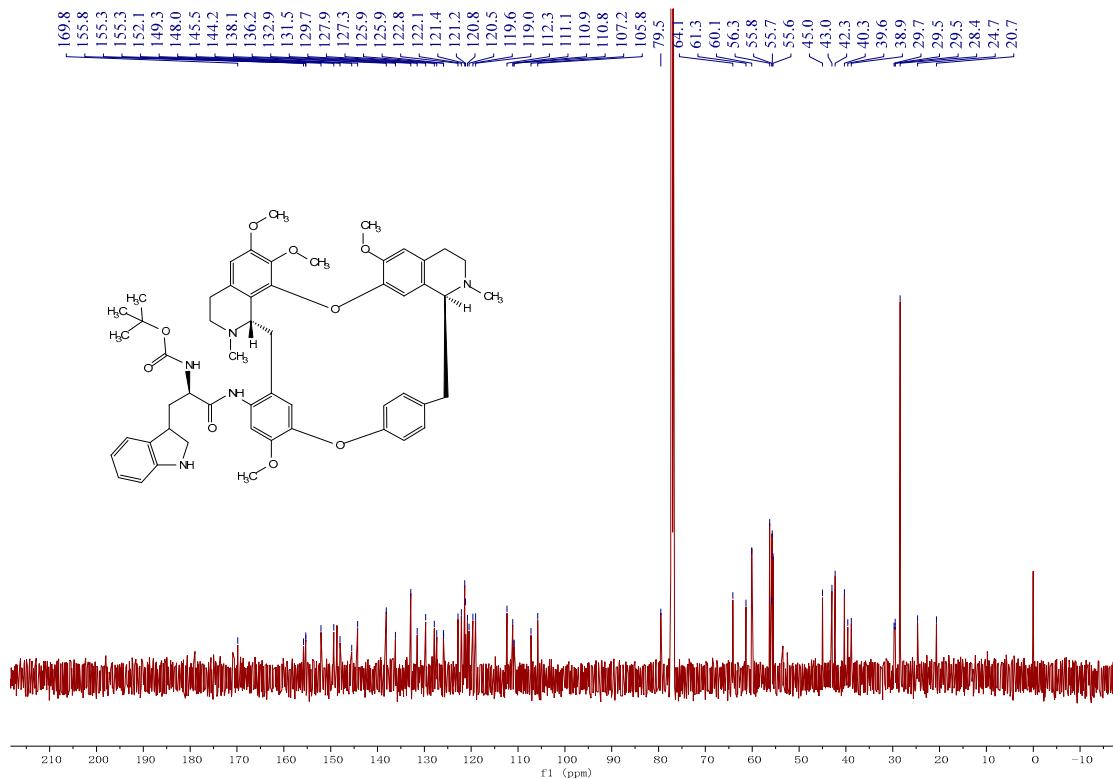
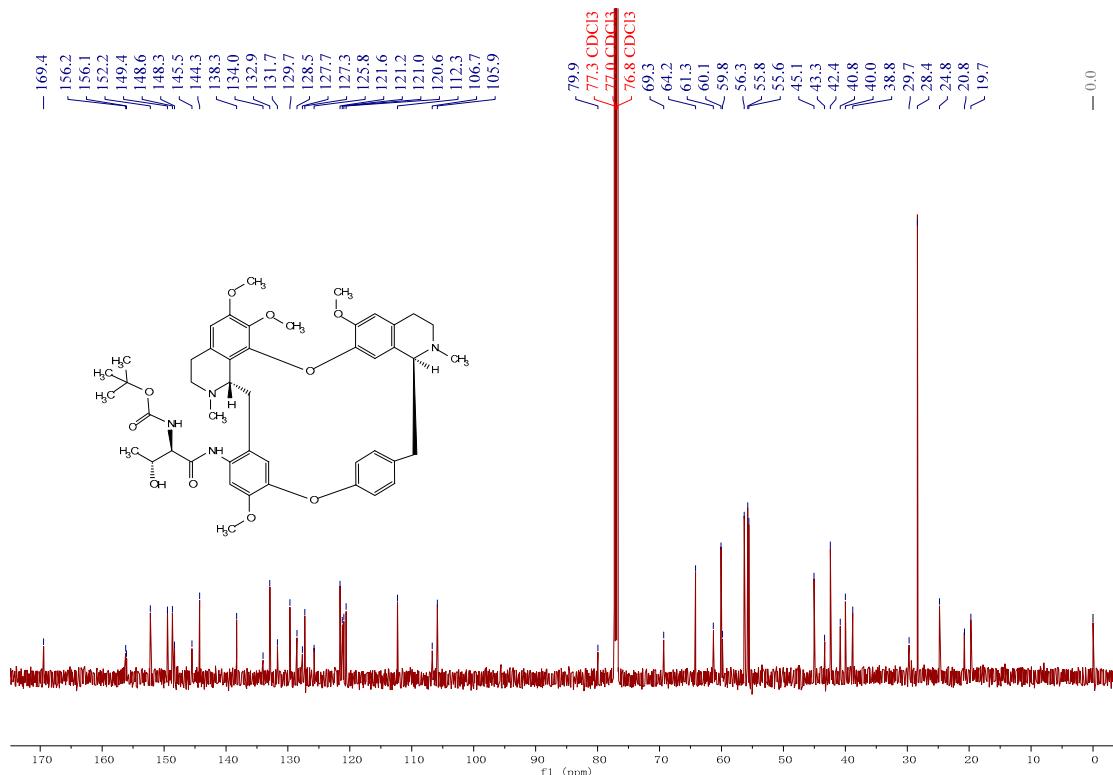


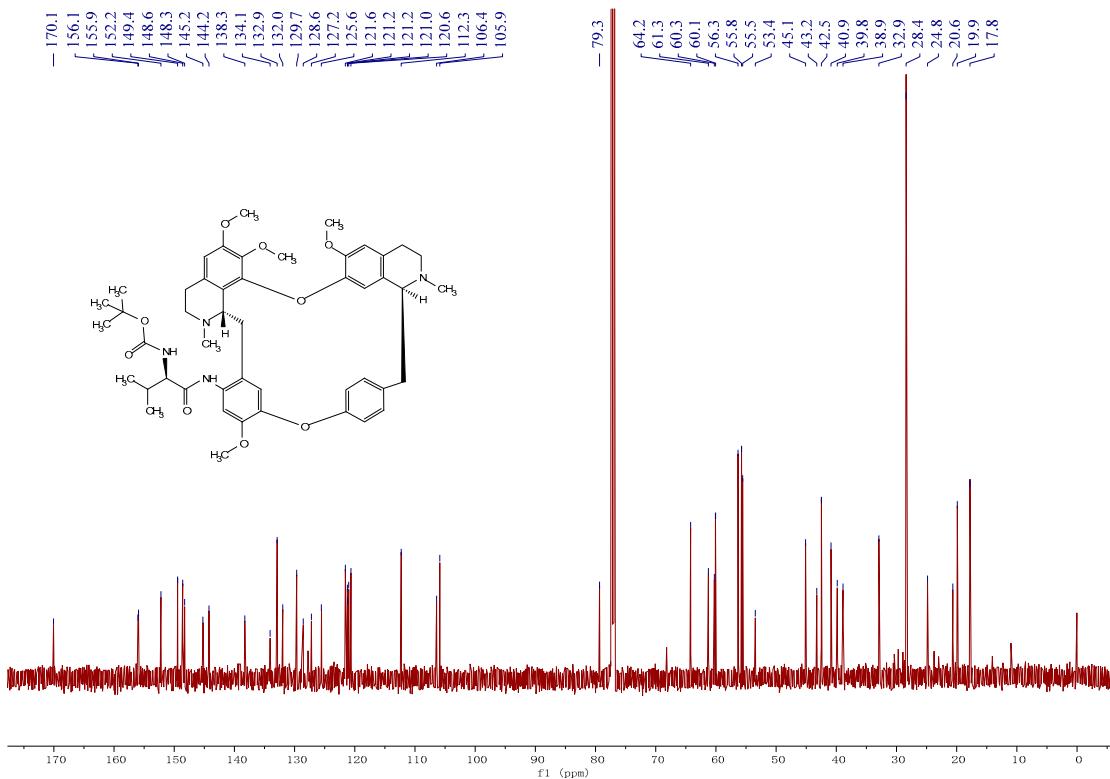
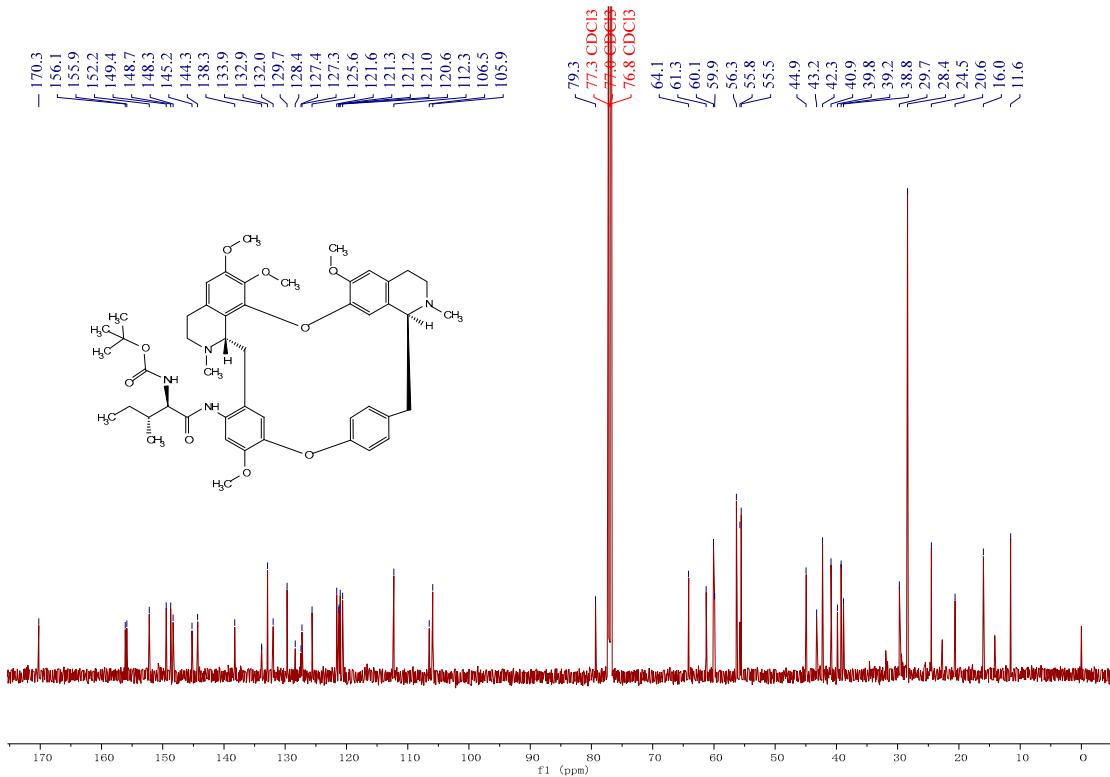
**Figure S29.**  $^{13}\text{C}$ -NMR Spectra of **1a** in  $\text{CDCl}_3$ .

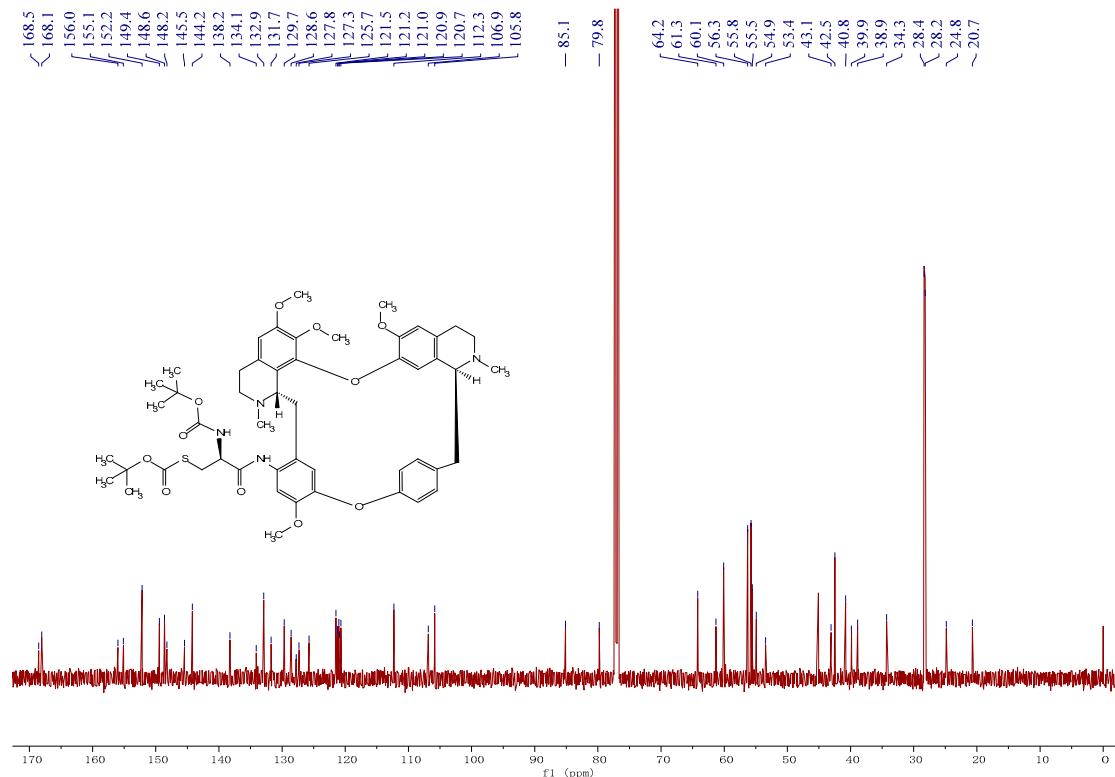
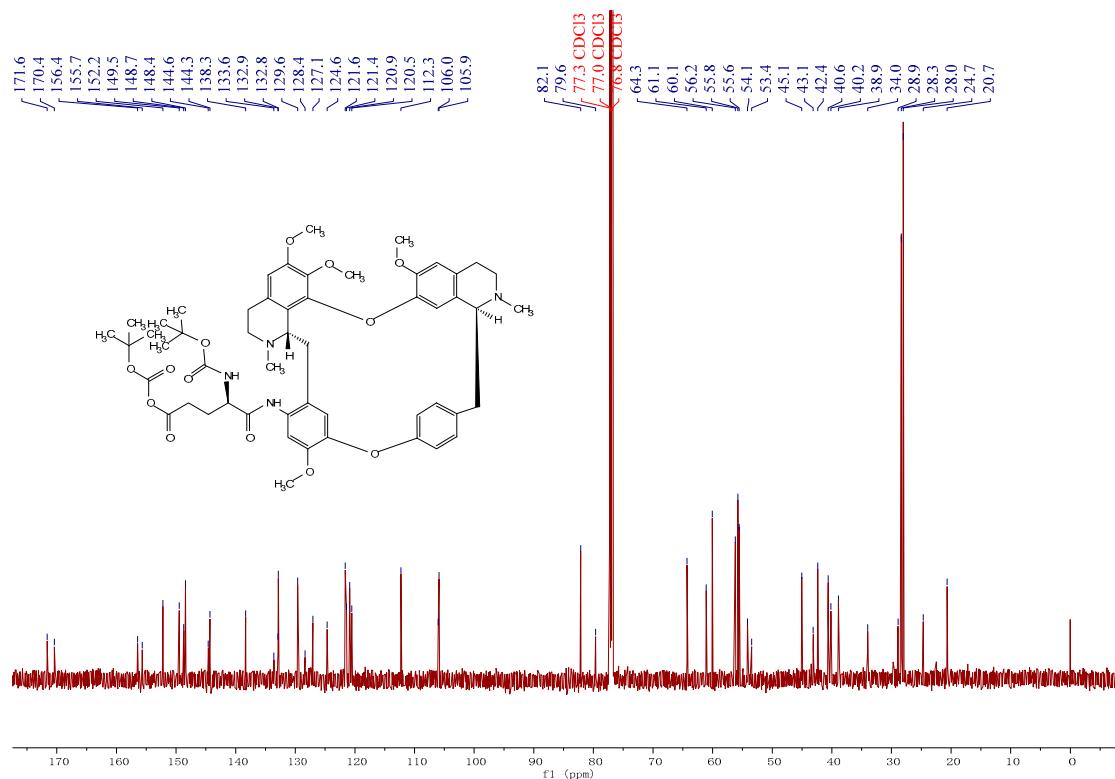


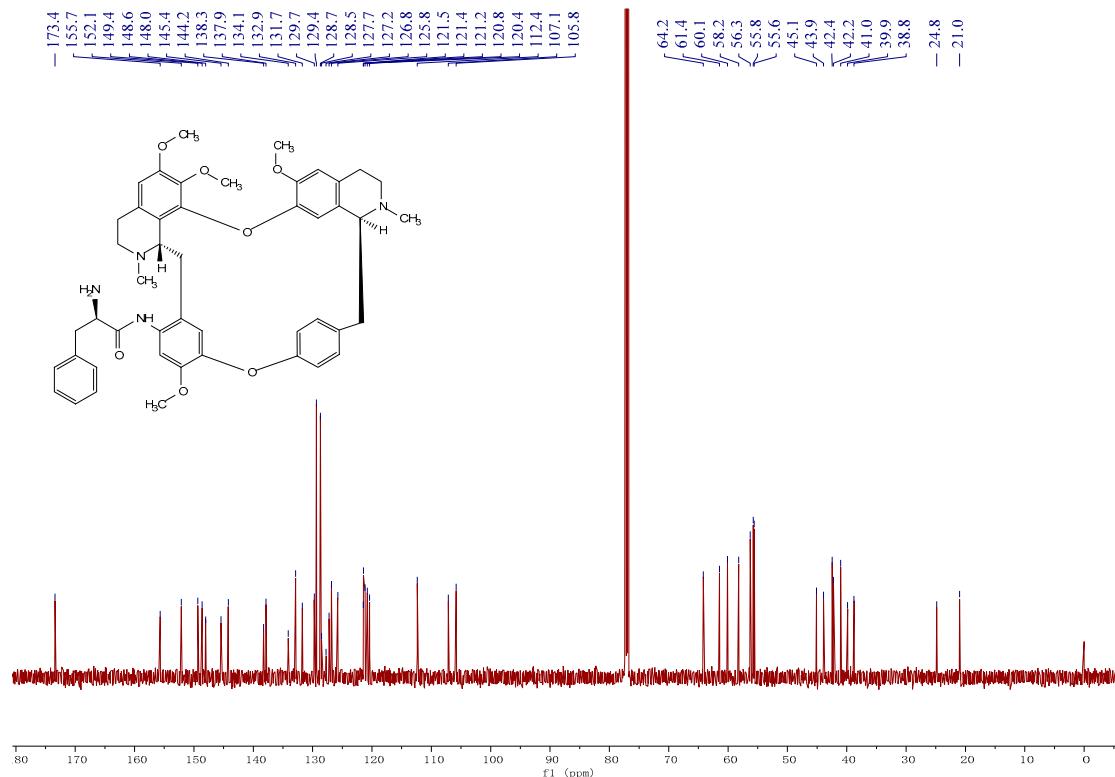
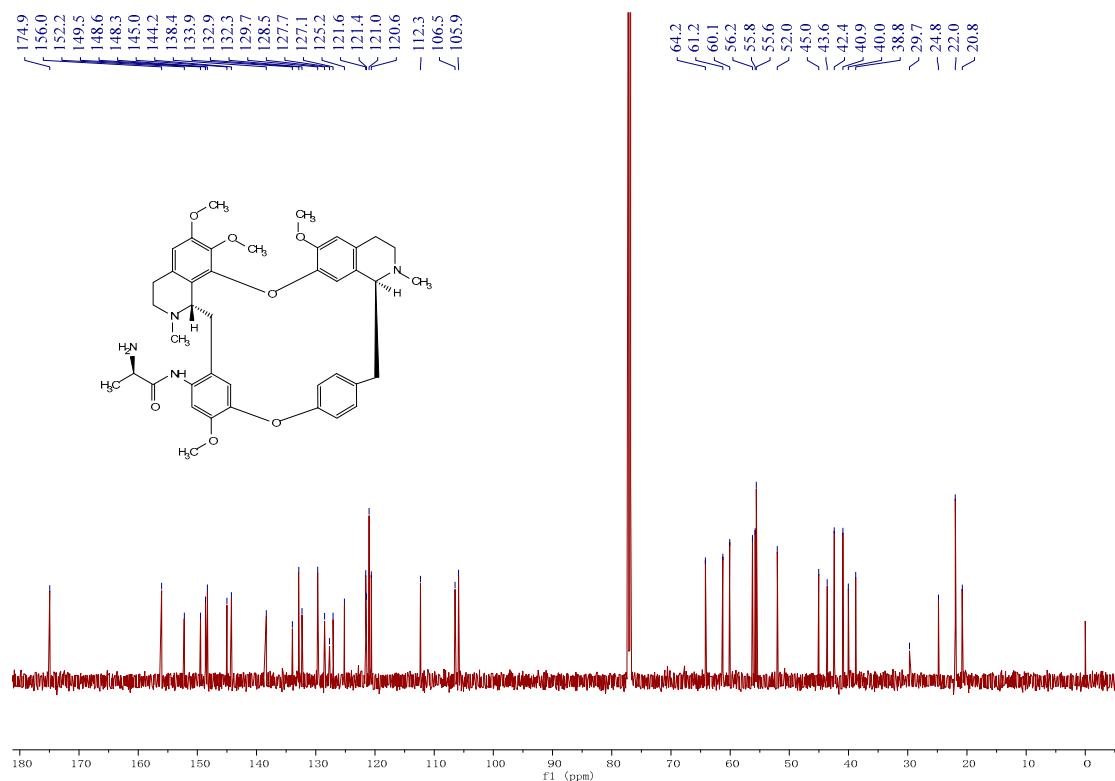
**Figure S30.**  $^{13}\text{C}$ -NMR Spectra of **1b** in  $\text{CDCl}_3$ .

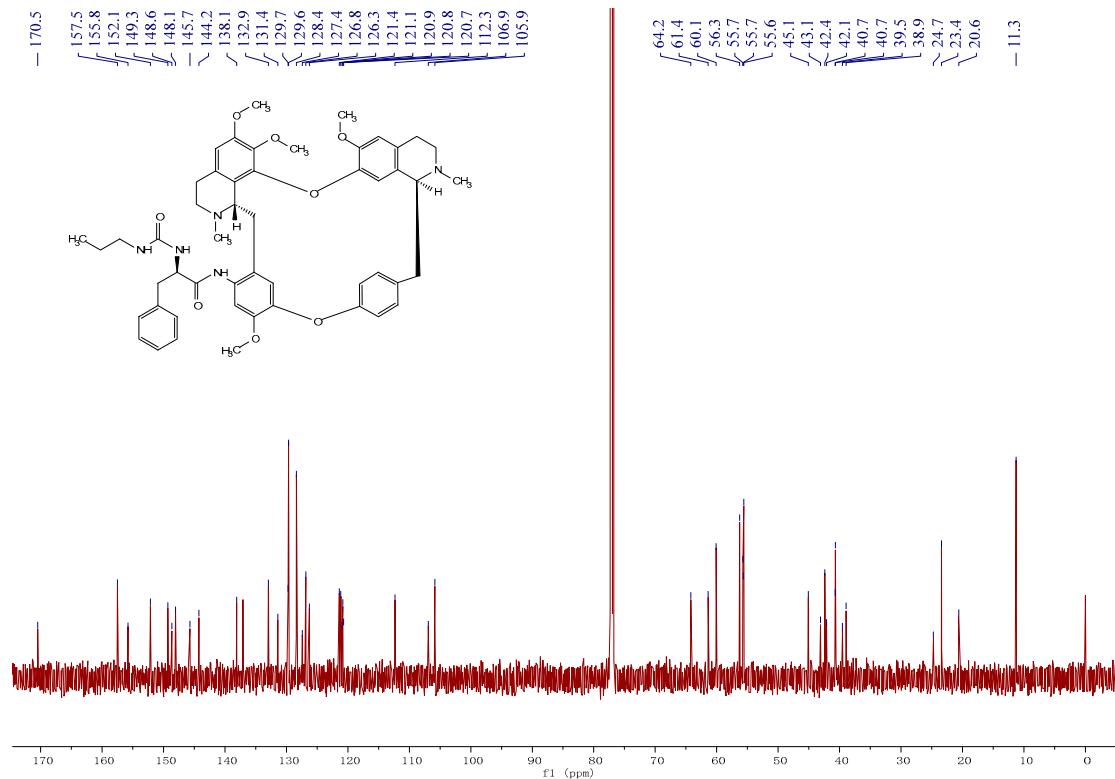
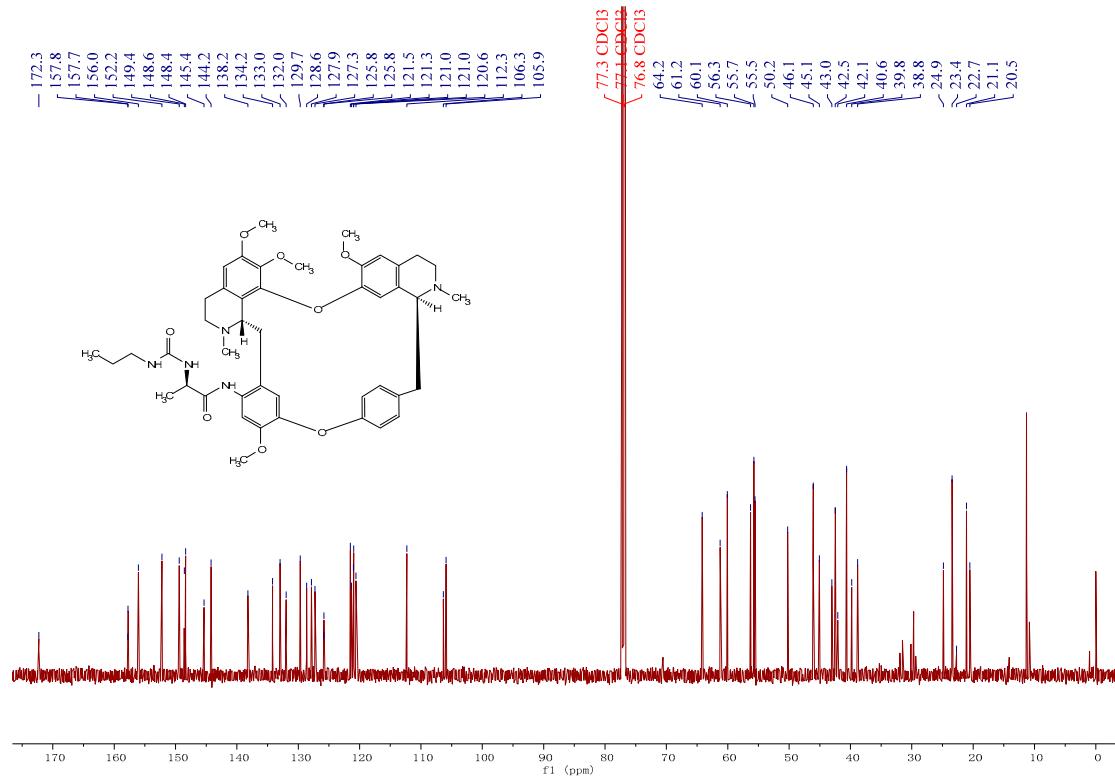
**Figure S31.** <sup>13</sup>C-NMR Spectra of **1c** in  $\text{CDCl}_3$ .**Figure S32.** <sup>13</sup>C-NMR Spectra of **1d** in  $\text{CDCl}_3$ .

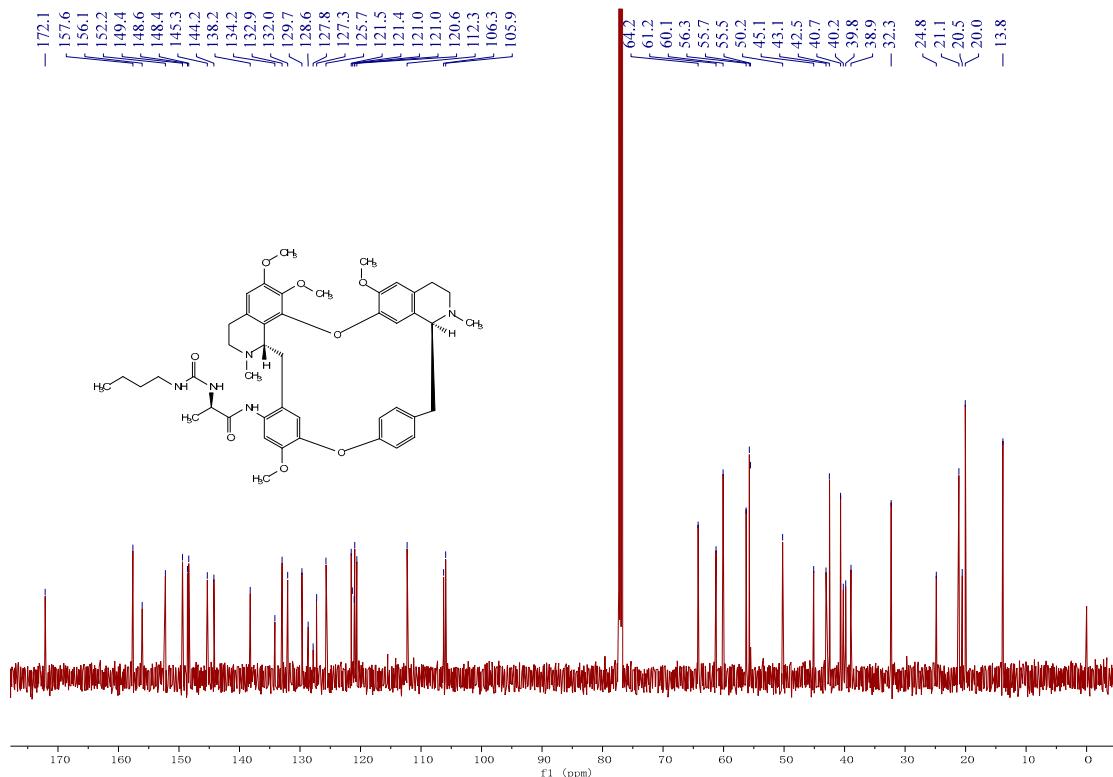
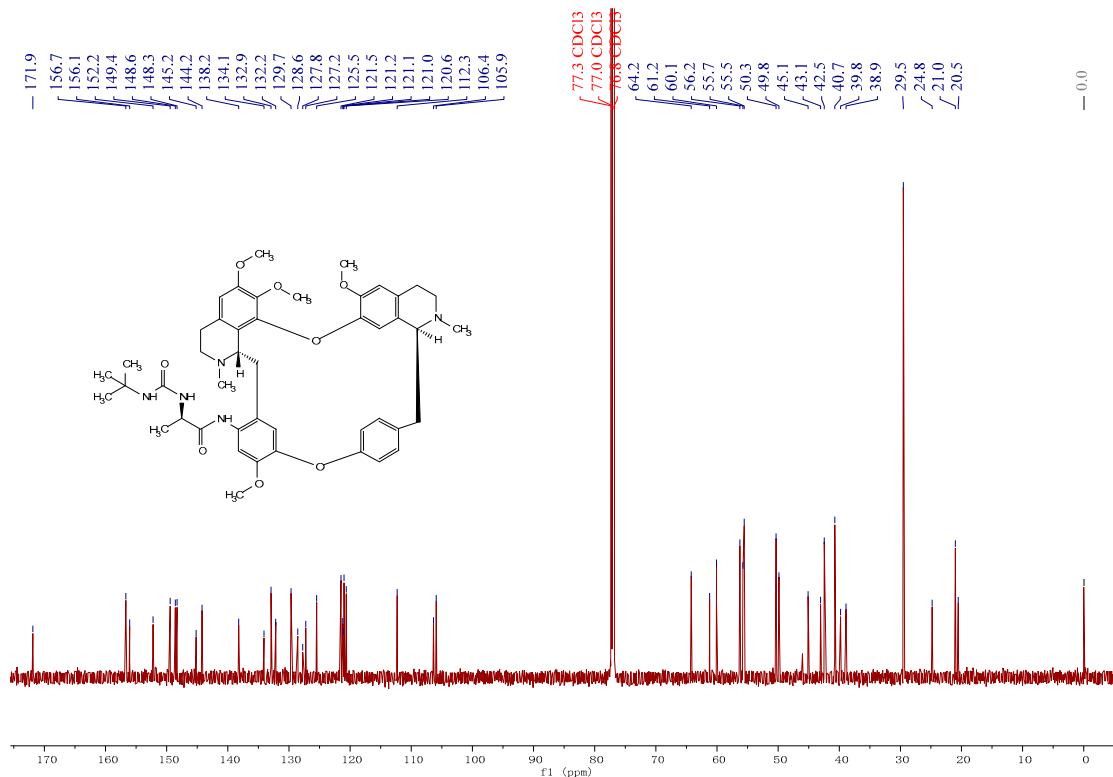
**Figure S33.**  $^{13}\text{C}$ -NMR Spectra of **1e** in  $\text{CDCl}_3$ .**Figure S34.**  $^{13}\text{C}$ -NMR Spectra of **1f** in  $\text{CDCl}_3$ .

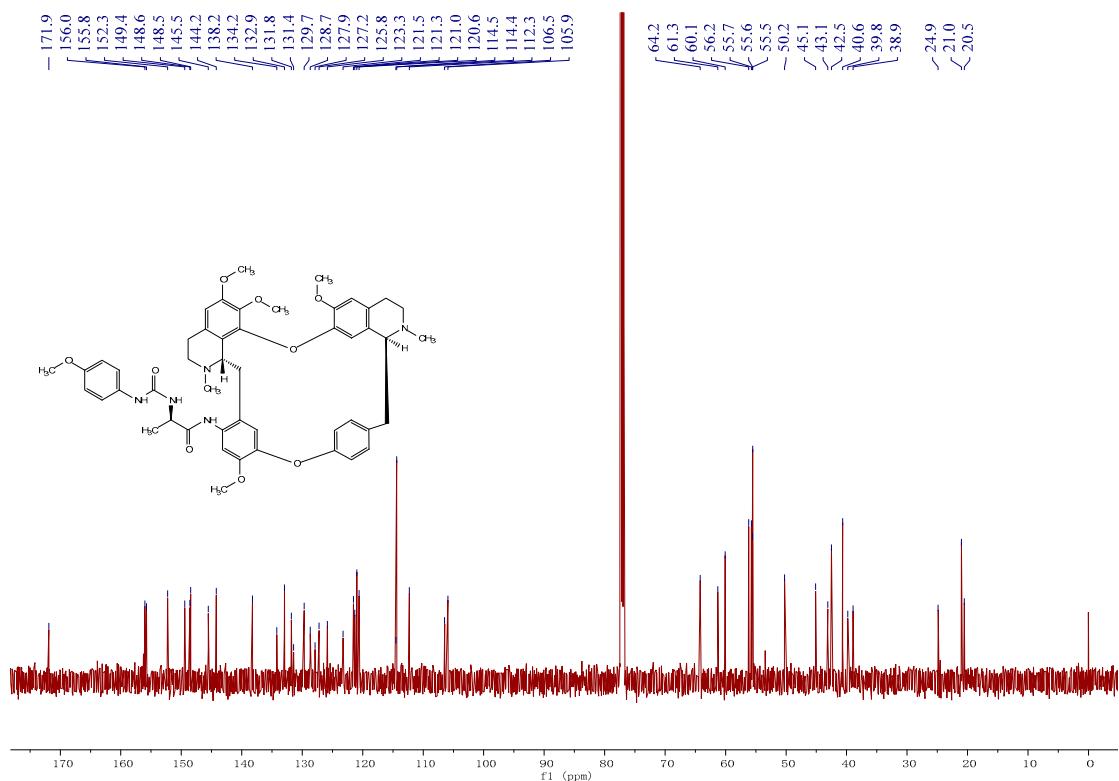
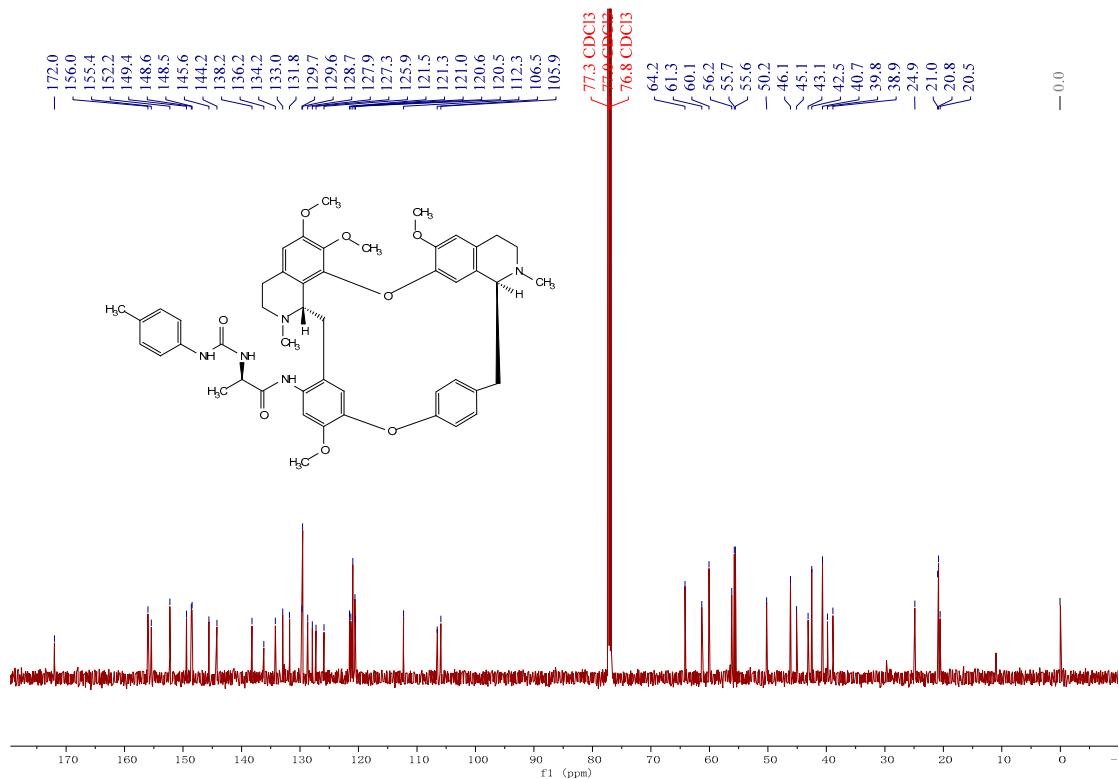
**Figure S35.**  $^{13}\text{C}$ -NMR Spectra of **1g** in  $\text{CDCl}_3$ .**Figure S36.**  $^{13}\text{C}$ -NMR Spectra of **1h** in  $\text{CDCl}_3$ .

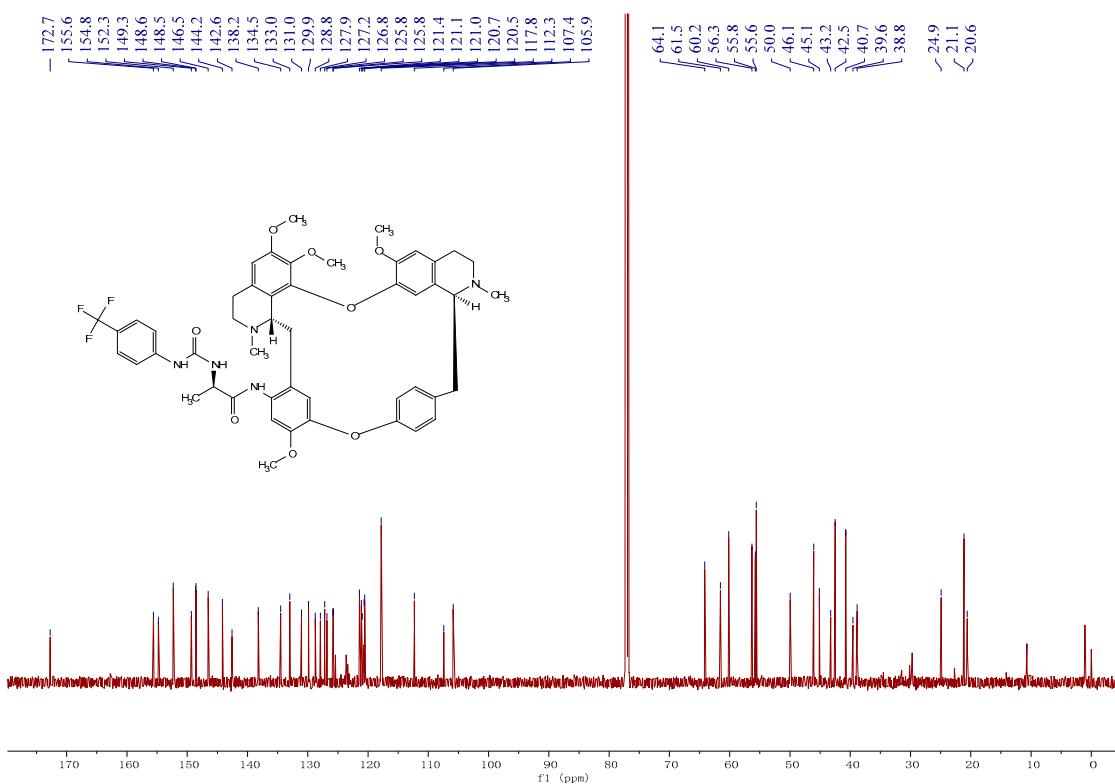
**Figure S37.**  $^{13}\text{C}$ -NMR Spectra of **1i** in  $\text{CDCl}_3$ .**Figure S38.**  $^{13}\text{C}$ -NMR Spectra of **1j** in  $\text{CDCl}_3$ .

**Figure S39.** <sup>13</sup>C-NMR Spectra of **1k** in  $\text{CDCl}_3$ .**Figure S40.** <sup>13</sup>C-NMR Spectra of **1l** in  $\text{CDCl}_3$ .

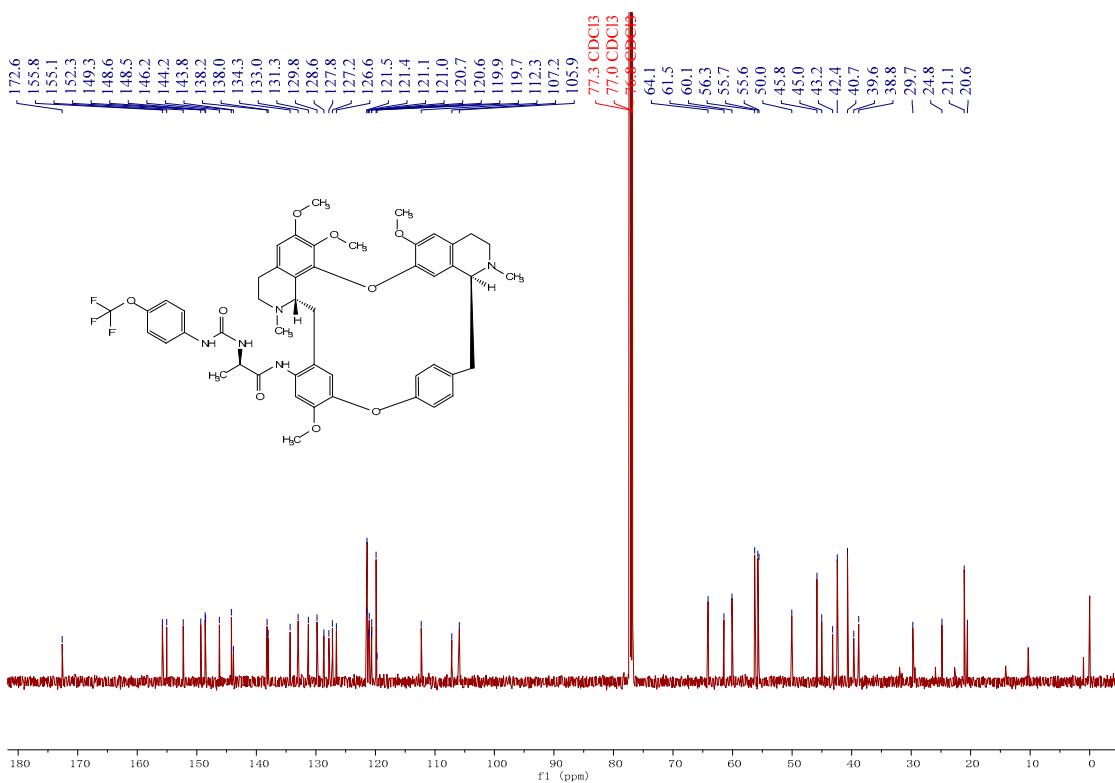
**Figure S41.**  $^{13}\text{C}$ -NMR Spectra of **2a** in  $\text{CDCl}_3$ .**Figure S42.**  $^{13}\text{C}$ -NMR Spectra of **3a** in  $\text{CDCl}_3$ .

**Figure S43.** <sup>13</sup>C-NMR Spectra of **3b** in CDCl<sub>3</sub>.**Figure S44.** <sup>13</sup>C-NMR Spectra of **3c** in CDCl<sub>3</sub>.

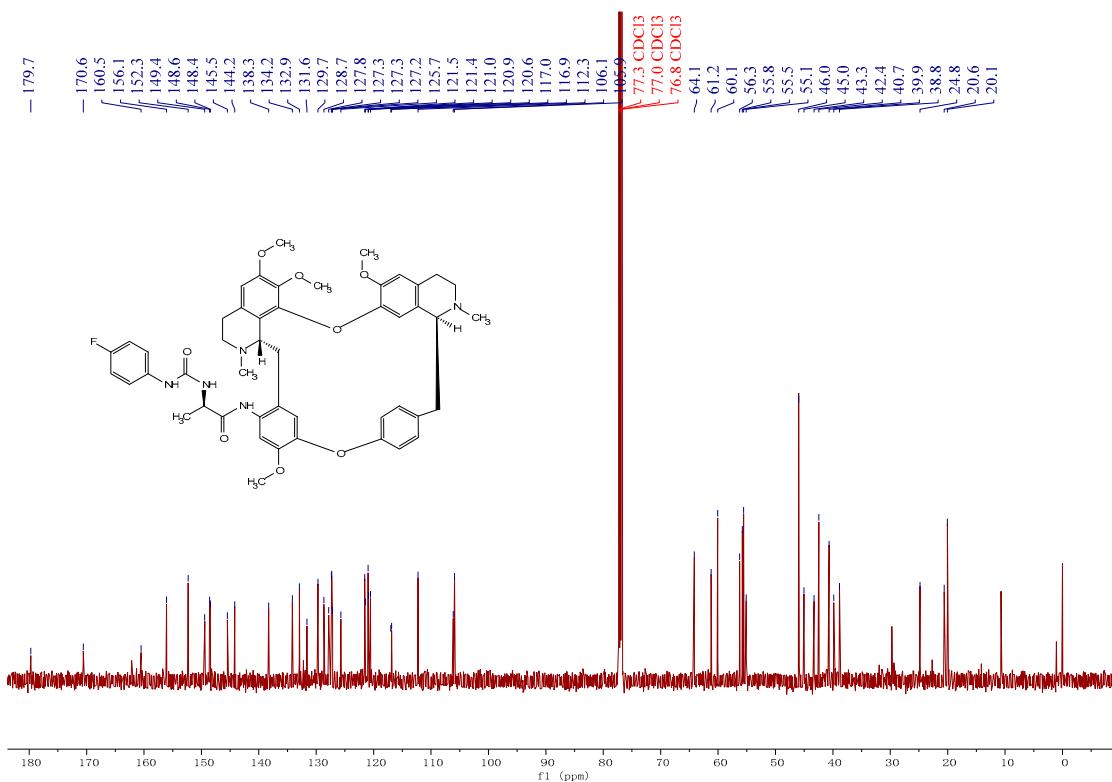
**Figure S46.**  $^{13}\text{C}$ -NMR Spectra of **3e** in  $\text{CDCl}_3$ .



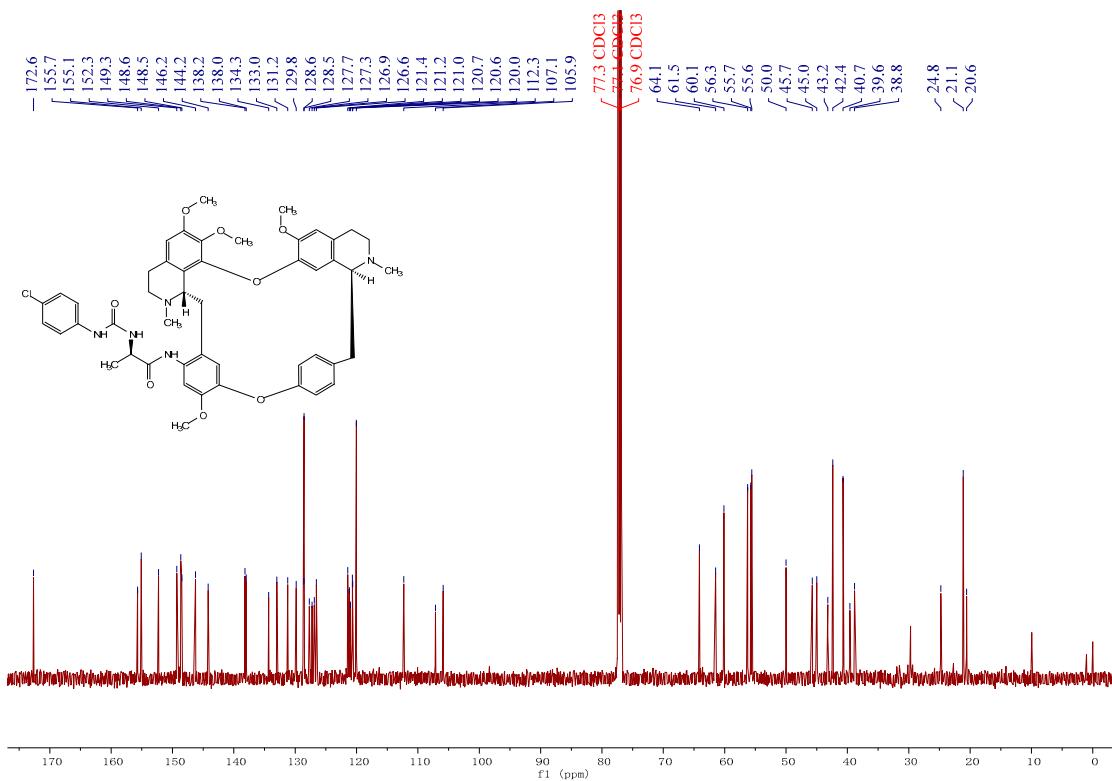
**Figure S47.**  $^{13}\text{C}$ -NMR Spectra of **3f** in  $\text{CDCl}_3$ .



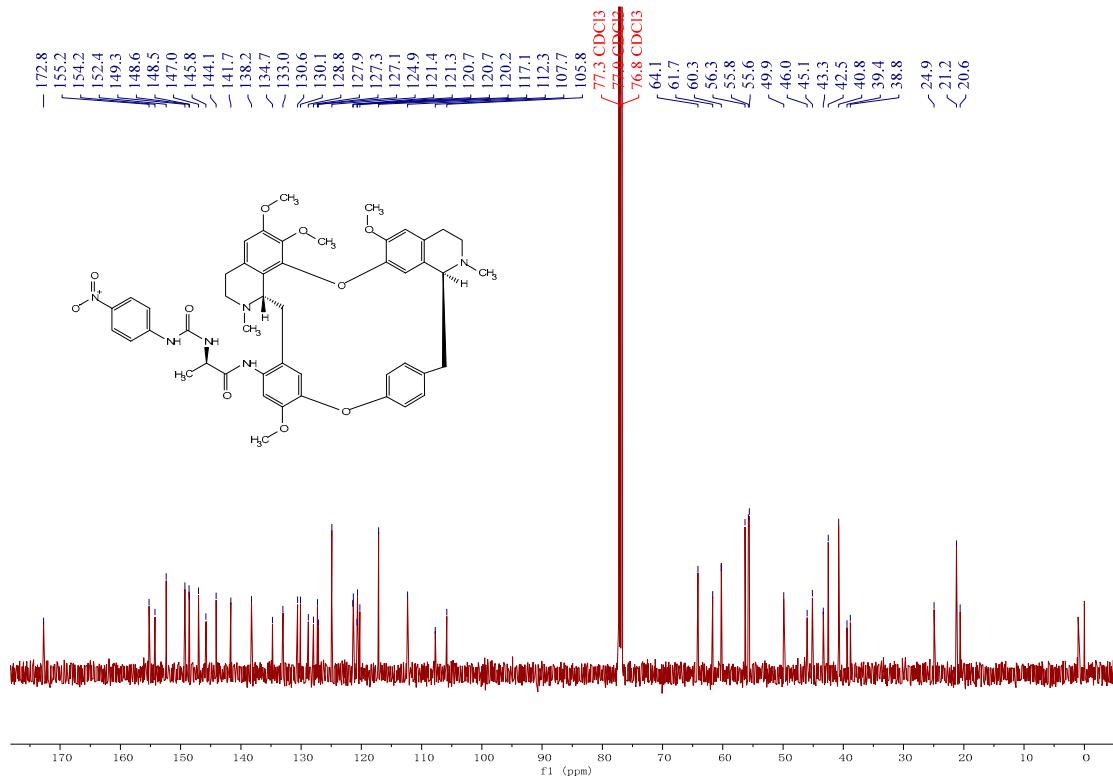
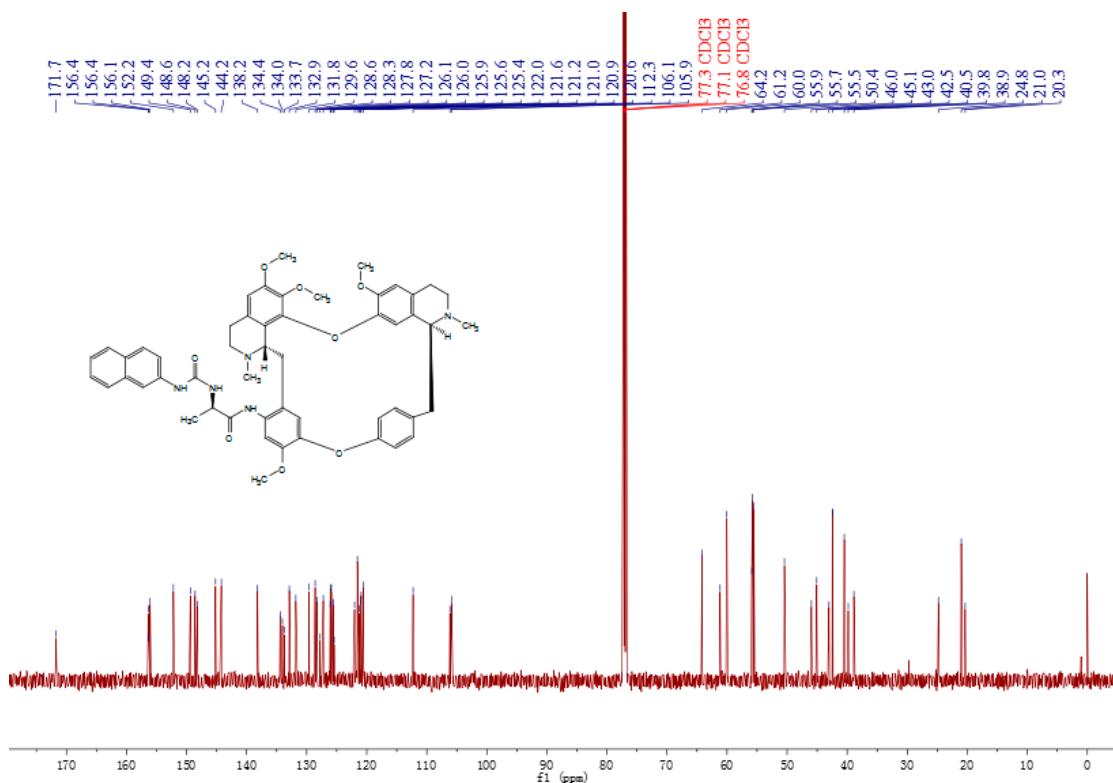
**Figure S48.**  $^{13}\text{C}$ -NMR Spectra of **3g** in  $\text{CDCl}_3$ .



**Figure S49.**  $^{13}\text{C}$ -NMR Spectra of **3h** in  $\text{CDCl}_3$ .



**Figure S50.**  $^{13}\text{C}$ -NMR Spectra of **3i** in  $\text{CDCl}_3$ .

**Figure S51.** <sup>13</sup>C-NMR Spectra of **3j** in CDCl<sub>3</sub>.**Figure S52.** <sup>13</sup>C-NMR Spectra of **3k** in CDCl<sub>3</sub>.