

# Synthesis of novel methyl 3-(hetero)arylthieno[3,2-*b*]pyridine-2-carboxylates and antitumor activity evaluation: studies *in vitro* and *in ovo* grafts of Chick Chorioallantoic Membrane (CAM) with a triple negative breast cancer cell line

Bruna R. Silva <sup>1,2,3,#</sup>, Rita Rebelo <sup>2,3,#</sup>, Juliana M. Rodrigues <sup>1</sup>, Cristina P. R. Xavier <sup>2,3\*</sup>, M. Helena Vasconcelos <sup>2,3,4</sup>, Maria-João R. P. Queiroz <sup>1\*</sup>

<sup>1</sup> Centre of Chemistry, University of Minho, 4710-057, Braga, Portugal; mjrqpq@quimica.uminho.pt (MJJQ); brunadansilva1967@gmail.com (BS); id7018@alunos.uminho.pt (JMR)

<sup>2</sup> Cancer Drug Resistance Group, IPATIMUP - Institute of Molecular Pathology and Immunology, University of Porto, Portugal, 4200-135, Porto, Portugal; rrebelo@ipatimup.pt (RR); cristinax@ipatimup.pt (CPRX); hvasconcelos@ipatimup.pt (MHV)

<sup>3</sup> i3S - Instituto de Investigação e Inovação em Saúde, Universidade do Porto, 4200-135 Porto, Portugal,

<sup>4</sup> Department of Biological Sciences, FFUP - Faculty of Pharmacy of the University of Porto, Porto, Portugal;

\* Correspondence: mjrqpq@quimica.uminho.pt (MJJRPQ); cristinax@ipatimup.pt (CPRX)

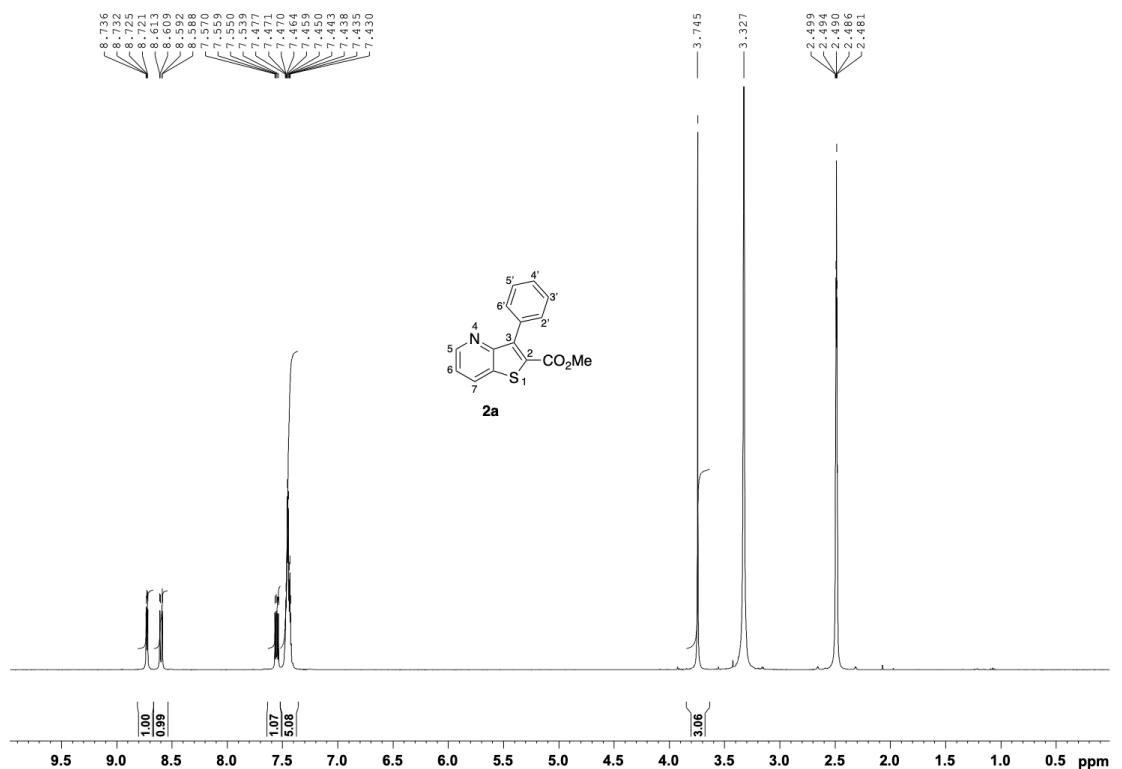
# These authors equally contributed to this work.

## Table of contents

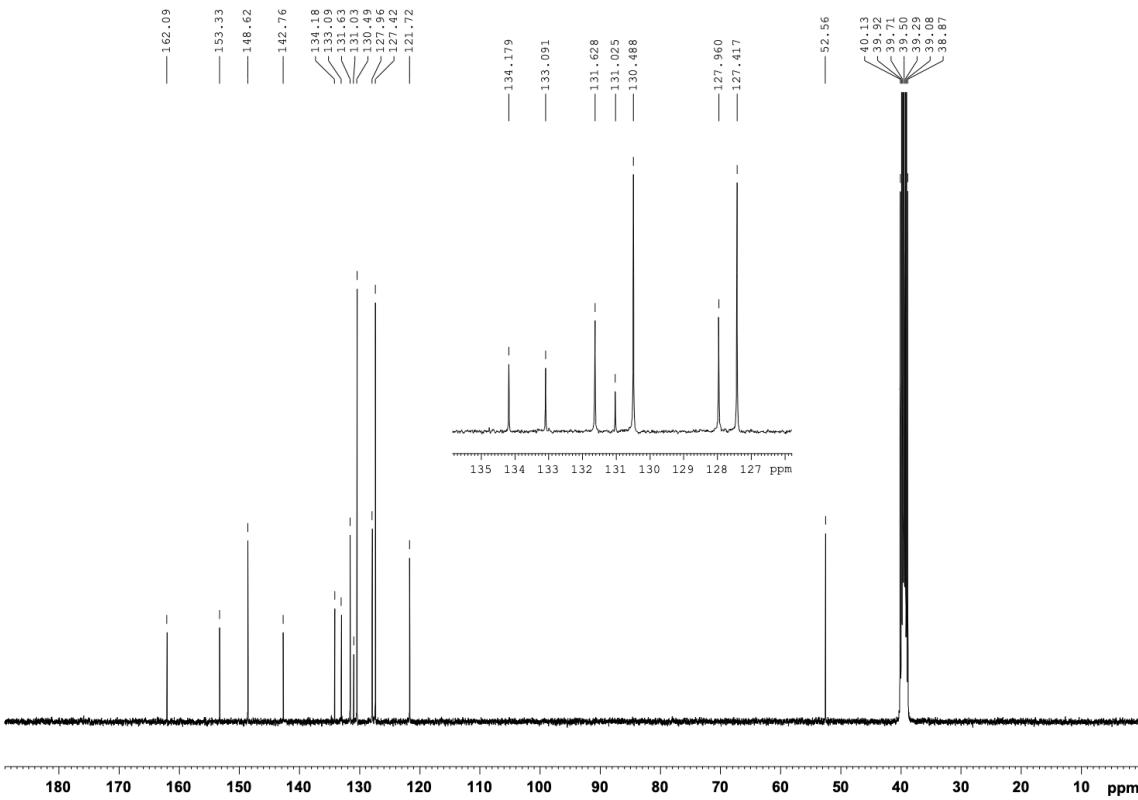
<sup>1</sup> H and <sup>13</sup> C NMR spectra of compound <b>2a</b> .....	S2
<sup>1</sup> H and <sup>13</sup> C NMR spectra of compound <b>2b</b> .....	S3
<sup>1</sup> H and <sup>13</sup> C NMR spectra of compound <b>2c</b> .....	S4
<sup>1</sup> H, <sup>13</sup> C NMR and <sup>19</sup> F spectra of compound <b>2d</b> .....	S5
<sup>1</sup> H and <sup>13</sup> C NMR spectra of compound <b>2e</b> .....	S7
<sup>1</sup> H and <sup>13</sup> C NMR spectra of compound <b>2f</b> .....	S8
<sup>1</sup> H and <sup>13</sup> C NMR spectra of compound <b>2g</b> .....	S9
<sup>1</sup> H and <sup>13</sup> C NMR spectra of compound <b>2h</b> .....	S10

### Methyl 3-phenylthieno[3,2-*b*]pyridine-2-carboxylate (2a)

<sup>1</sup>H NMR (DMSO-*d*<sub>6</sub>, 400 MHz) of compound **2a**

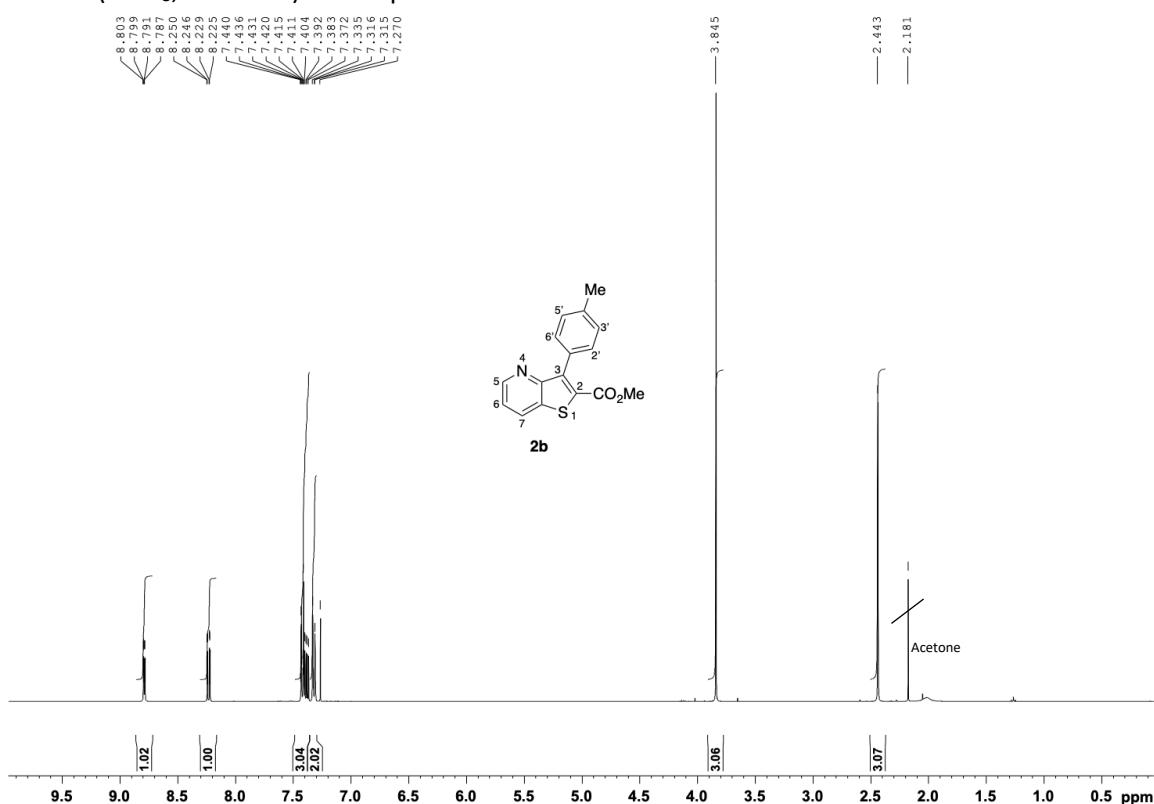


<sup>13</sup>C NMR (DMSO-*d*<sub>6</sub>, 100.6 MHz) of compound **2a**

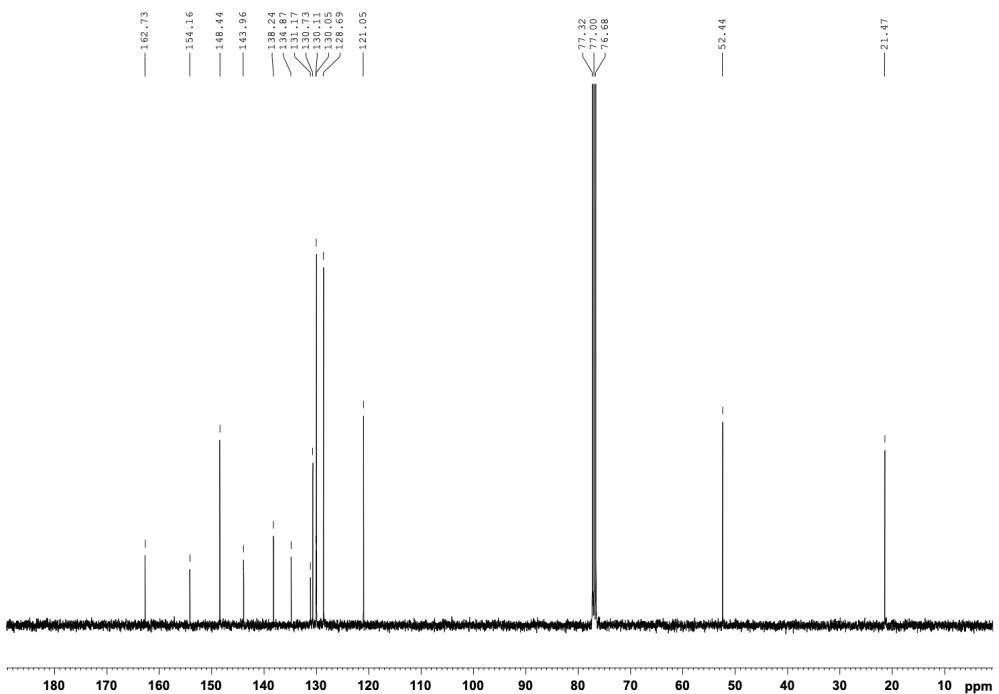


**Methyl 3-(*p*-tolyl)thieno[3,2-*b*]pyridine-2-carboxylate (**2b**)**

<sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz) of compound **2b**

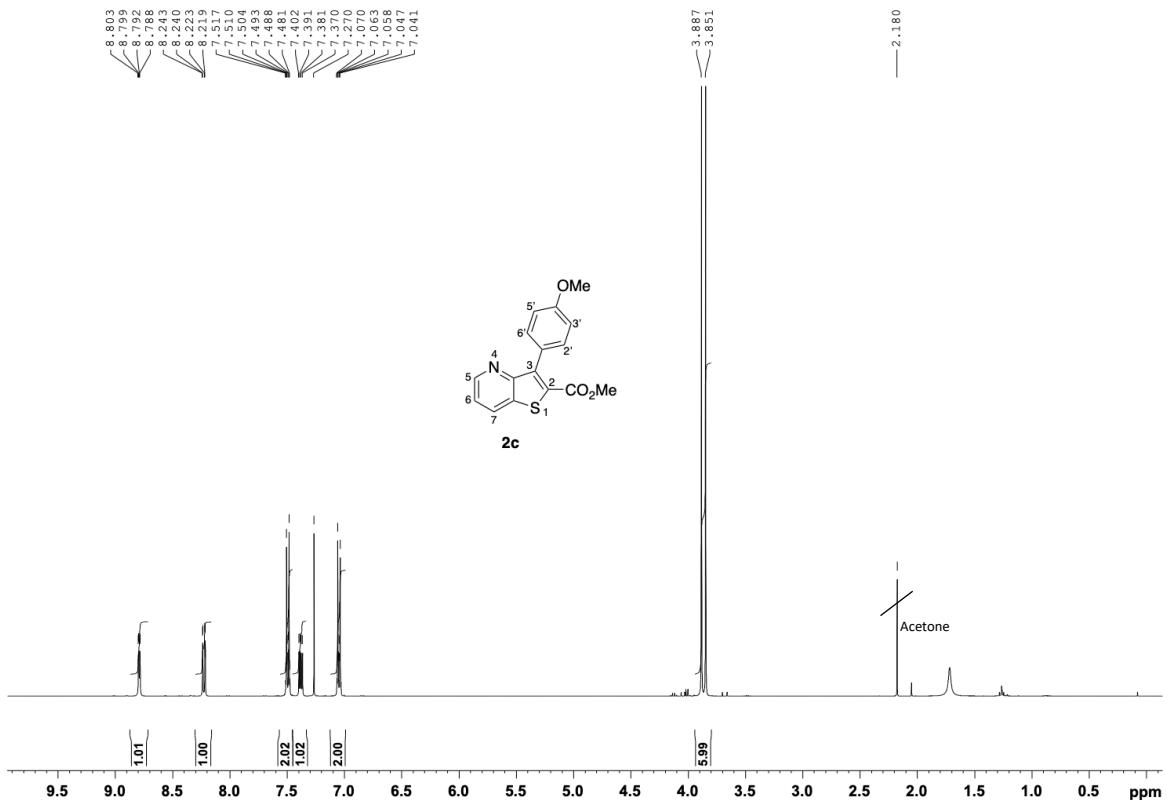


<sup>13</sup>C NMR (CDCl<sub>3</sub>, 100.6 MHz) of compound **2b**

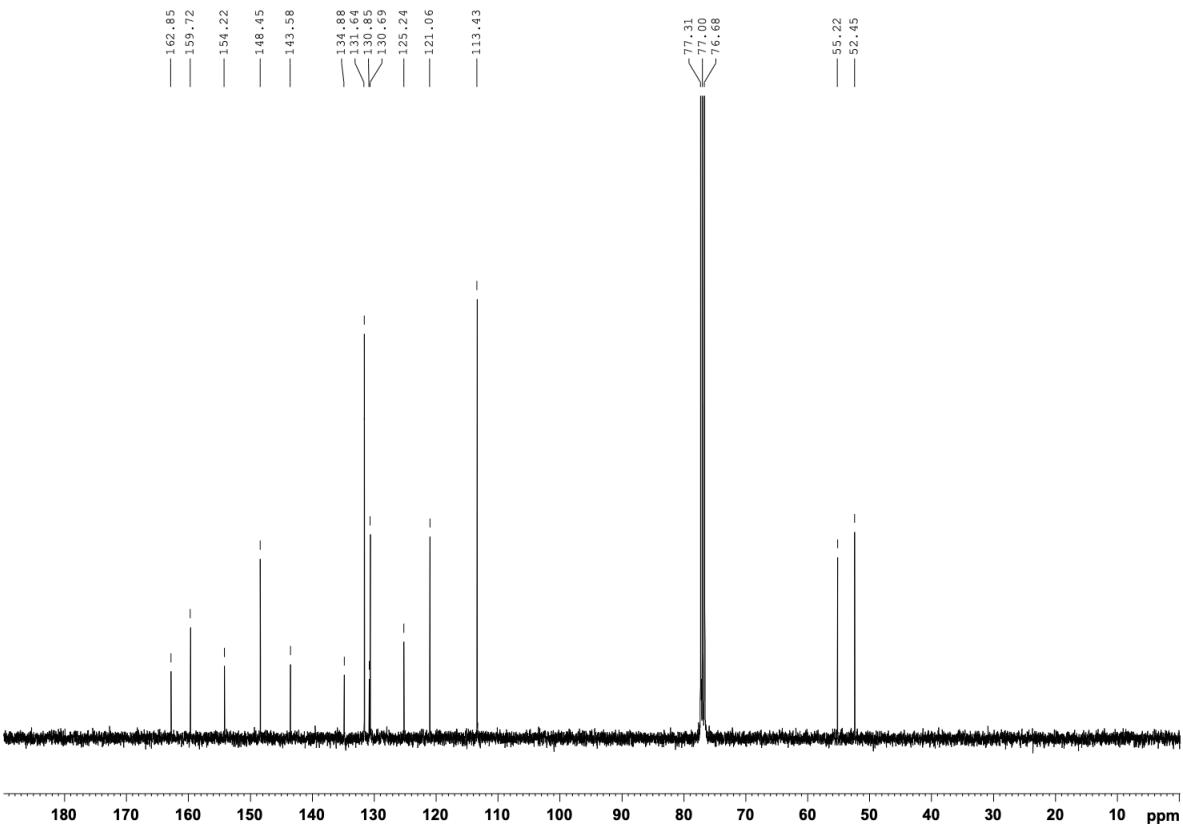


**Methyl 3-(4-methoxyphenyl)thieno[3,2-*b*]pyridine-2-carboxylate (**2c**)**

<sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz) of compound **2c**

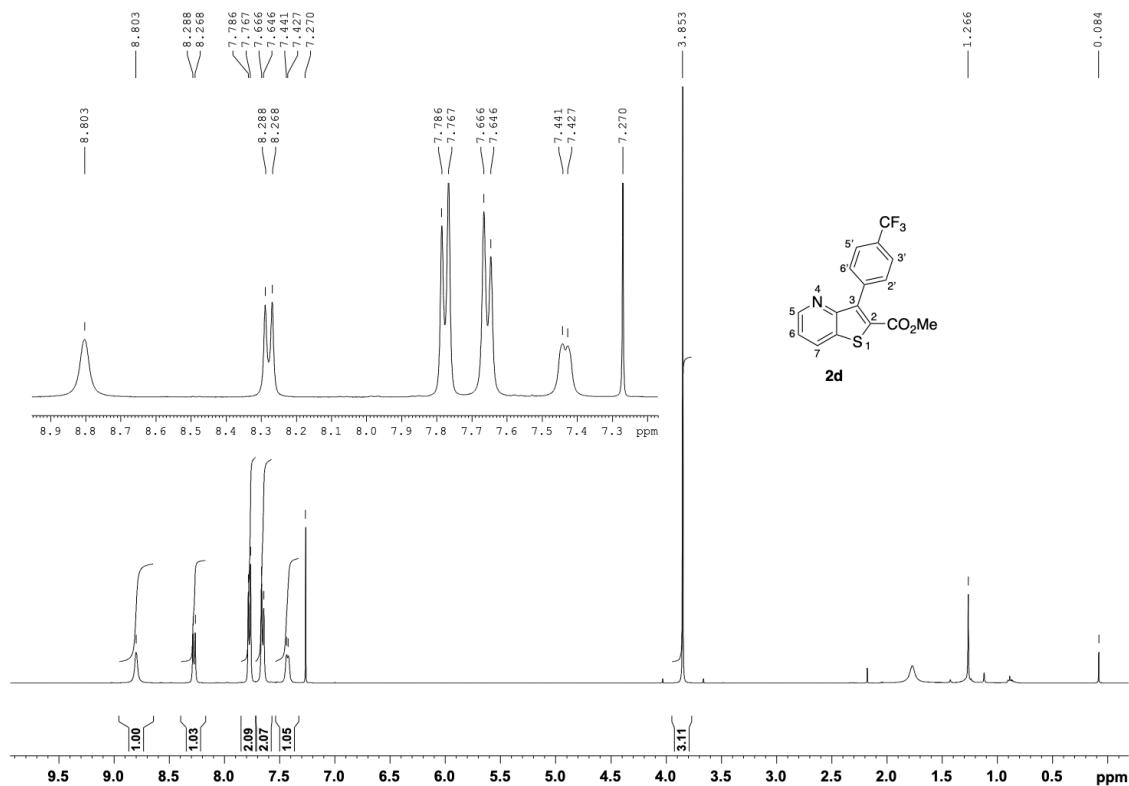


<sup>13</sup>C NMR (CDCl<sub>3</sub>, 100.6 MHz) of compound **2c**

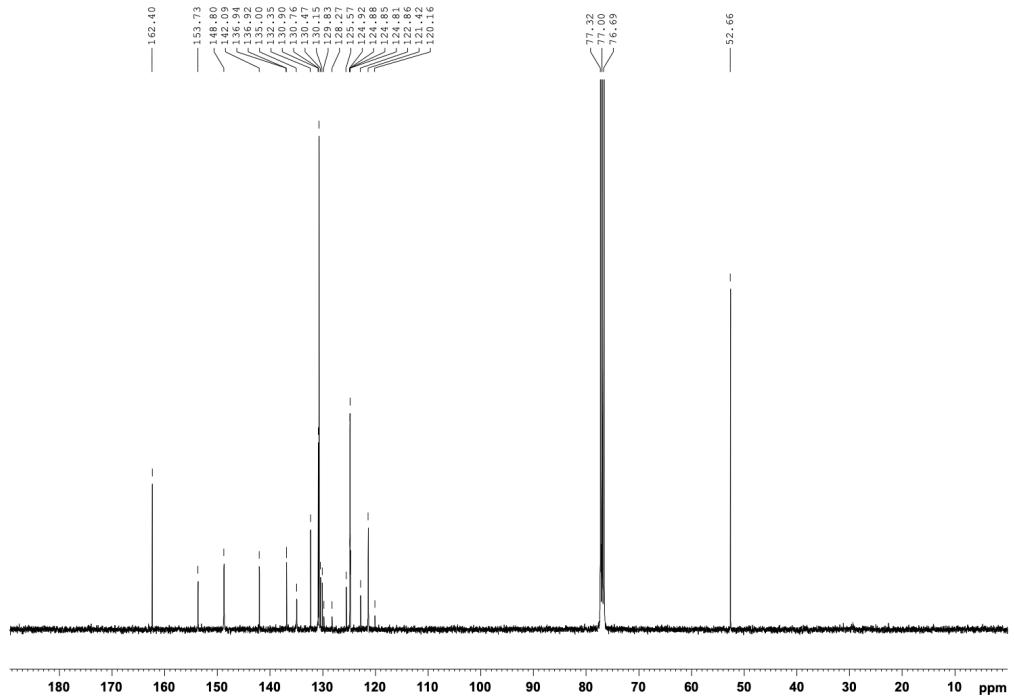


#### Methyl 3-[4-(trifluoromethyl)phenyl]thieno[3,2-*b*]pyridine-2-carboxylate (2d)

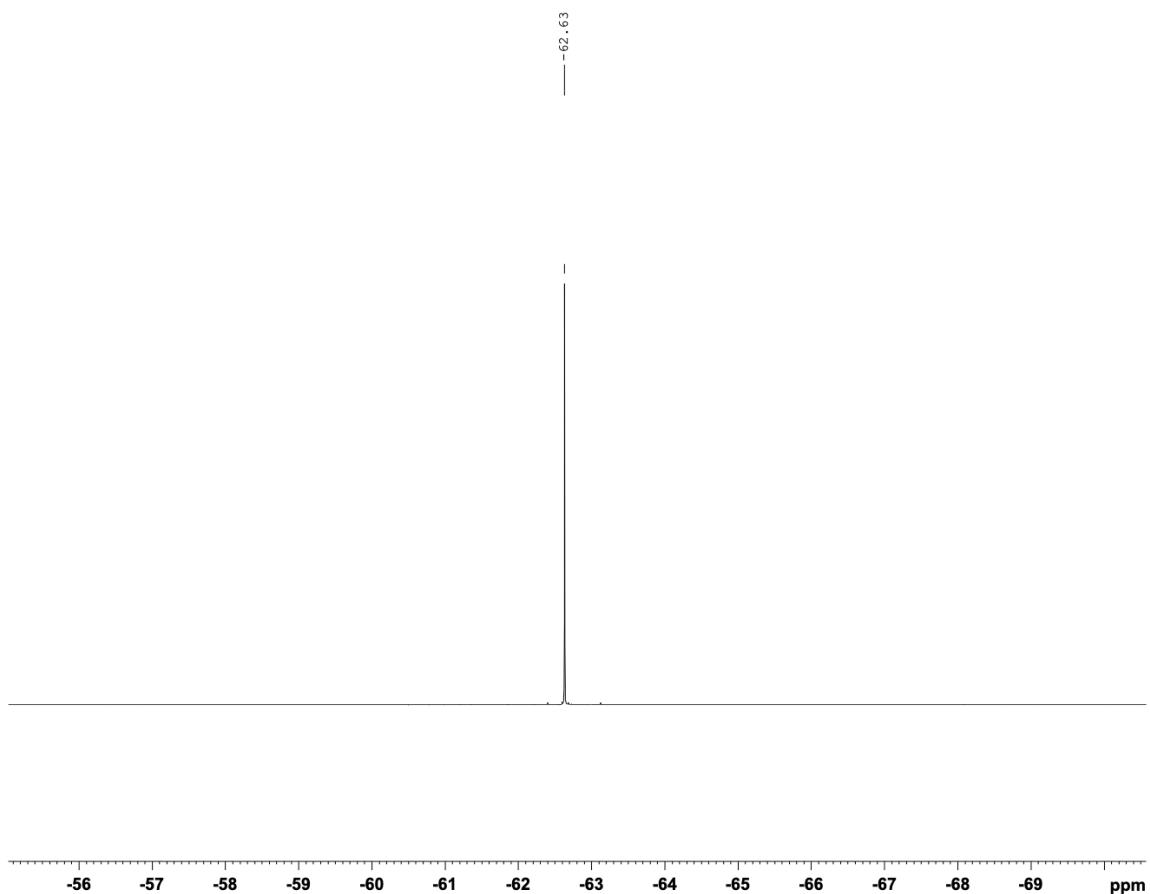
<sup>1</sup>H NMR ( $\text{CDCl}_3$ , 400 MHz) of compound **2d**



<sup>13</sup>C NMR ( $\text{CDCl}_3$ , 100.6 MHz) of compound **2d**

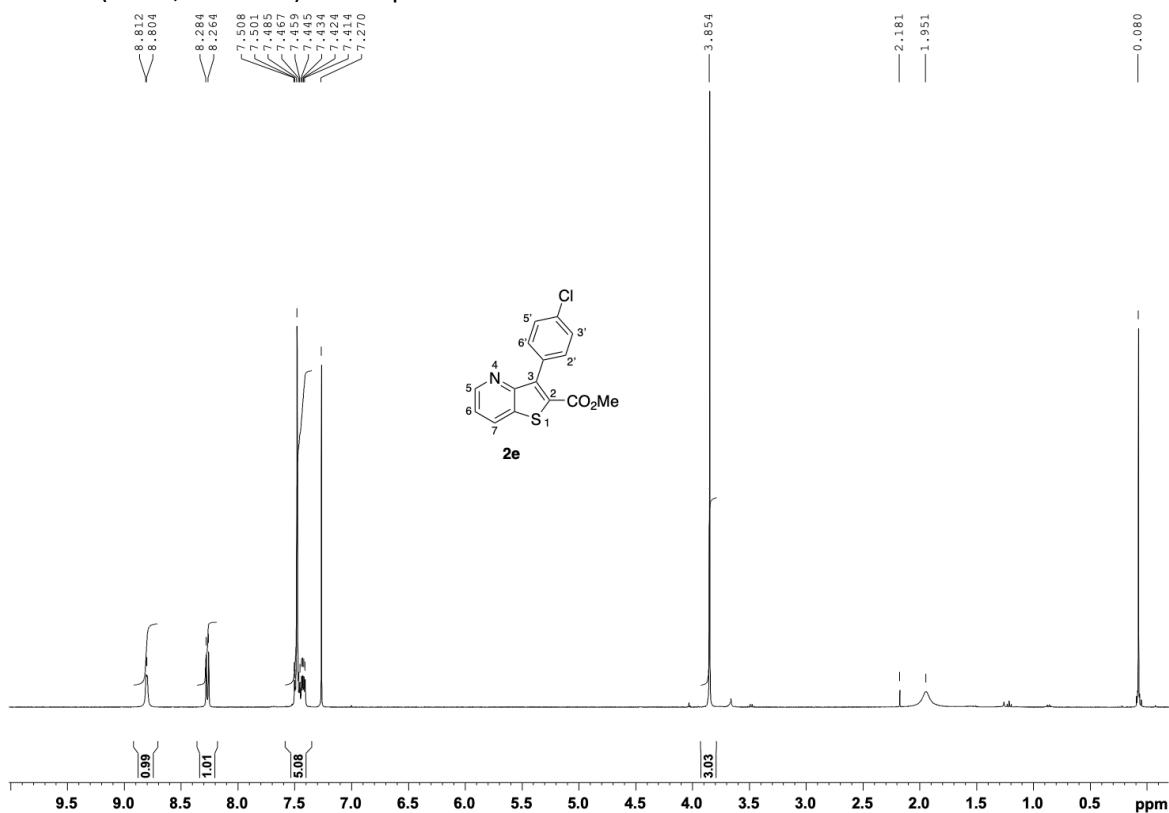


<sup>19</sup>F NMR ( $\text{CDCl}_3$ , 376.48 MHz) of compound **2d**

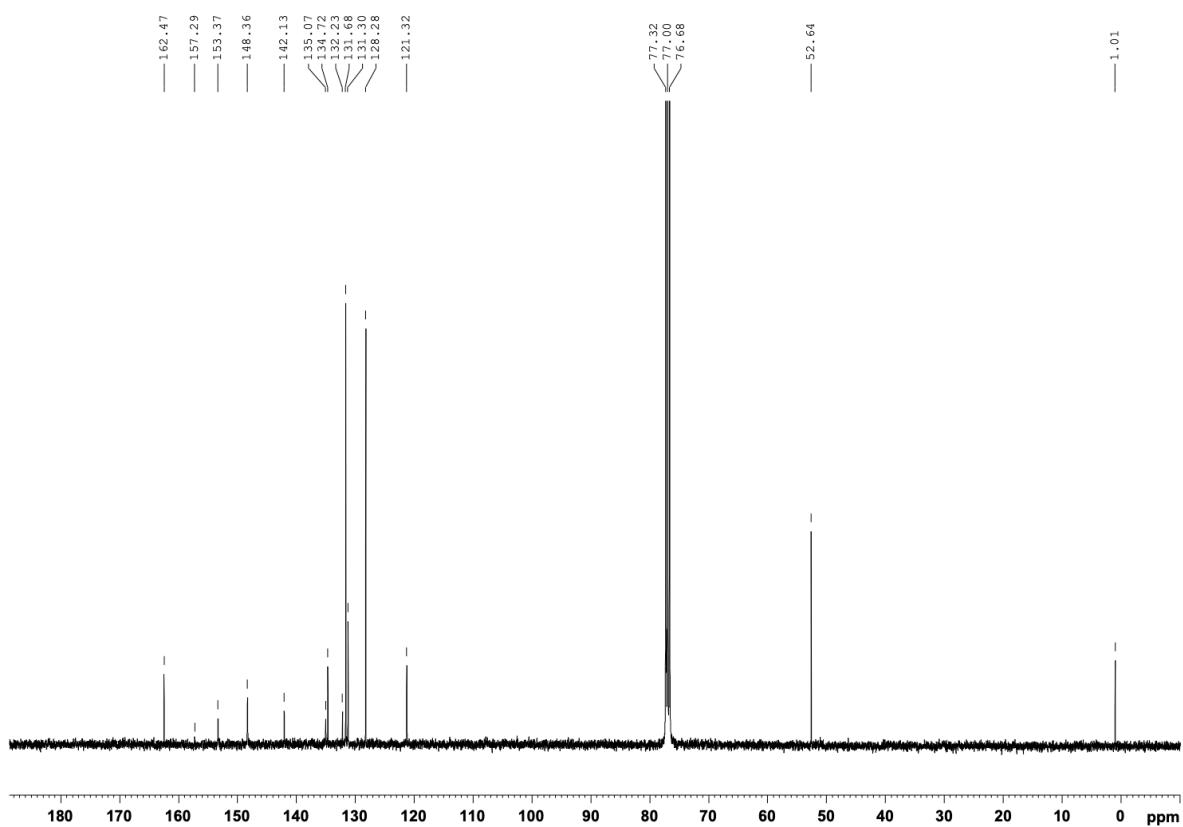


**Methyl 3-(4-chlorophenyl)thieno[3,2-*b*]pyridine-2-carboxylate (2e)**

<sup>1</sup>H NMR ( $\text{CDCl}_3$ , 400 MHz) of compound **2e**

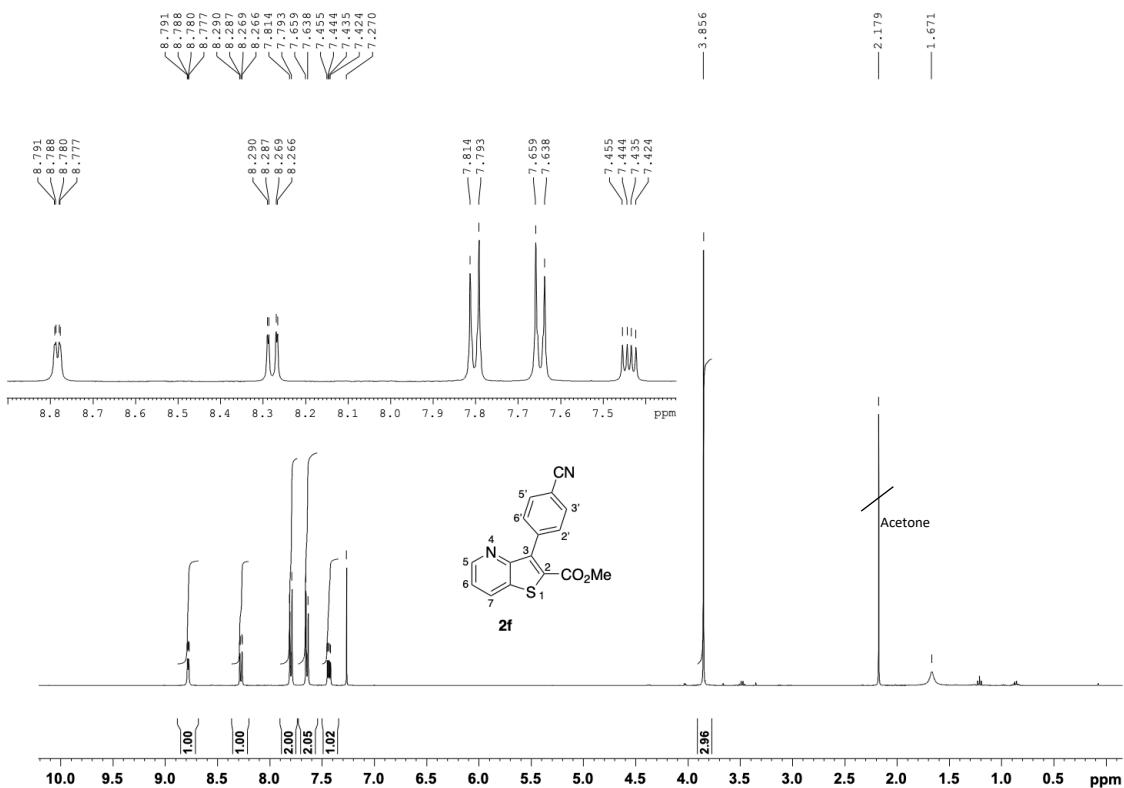


<sup>13</sup>C NMR ( $\text{CDCl}_3$ , 100.6 MHz) of compound **2e**

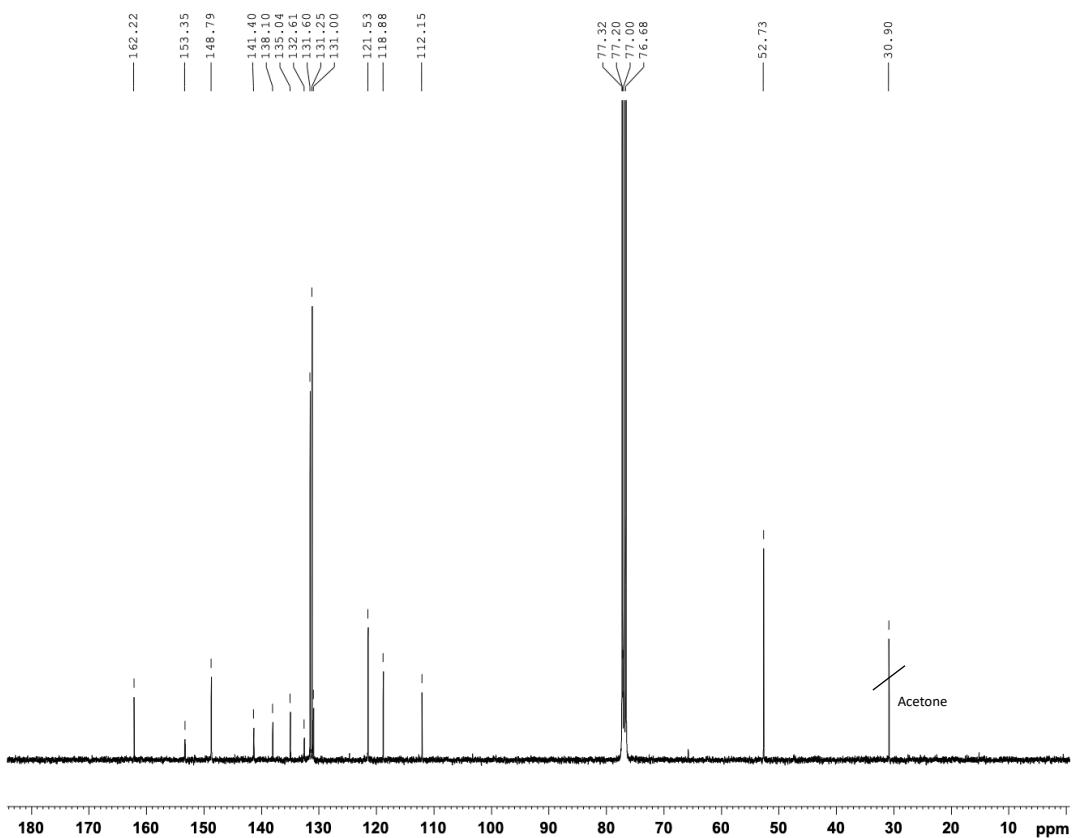


**Methyl 3-(4-cyanophenyl)thieno[3,2-*b*]pyridine-2-carboxylate (**2f**)**

<sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz) of compound **2f**

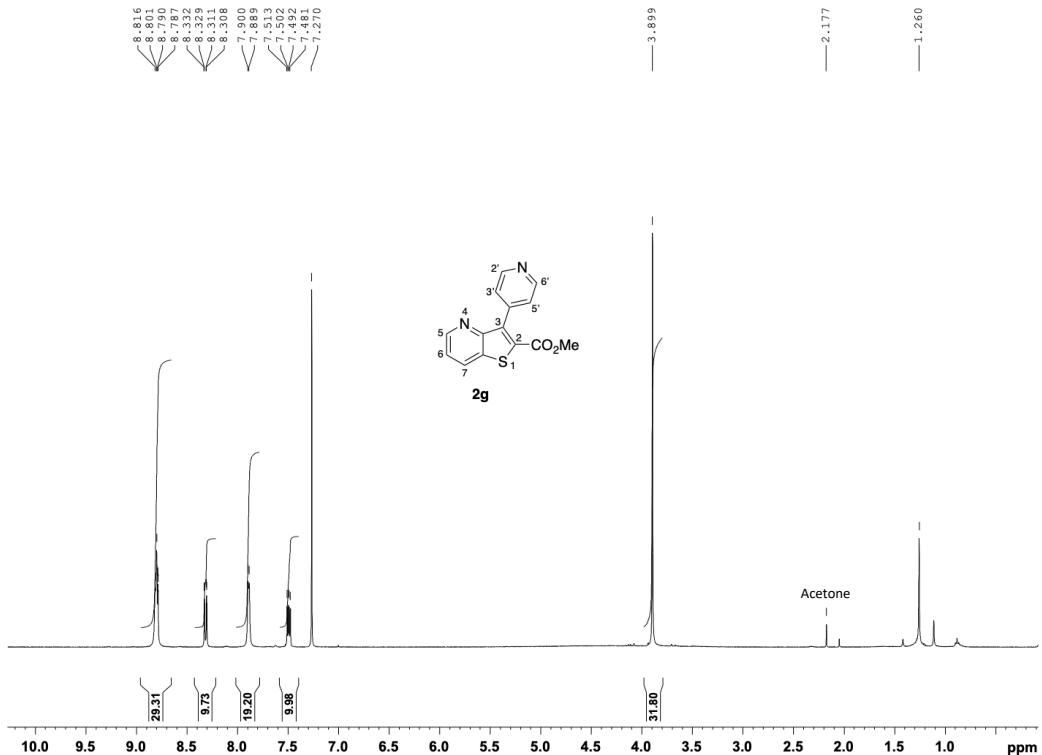


<sup>13</sup>C NMR (CDCl<sub>3</sub>, 100.6 MHz) of compound **2f**

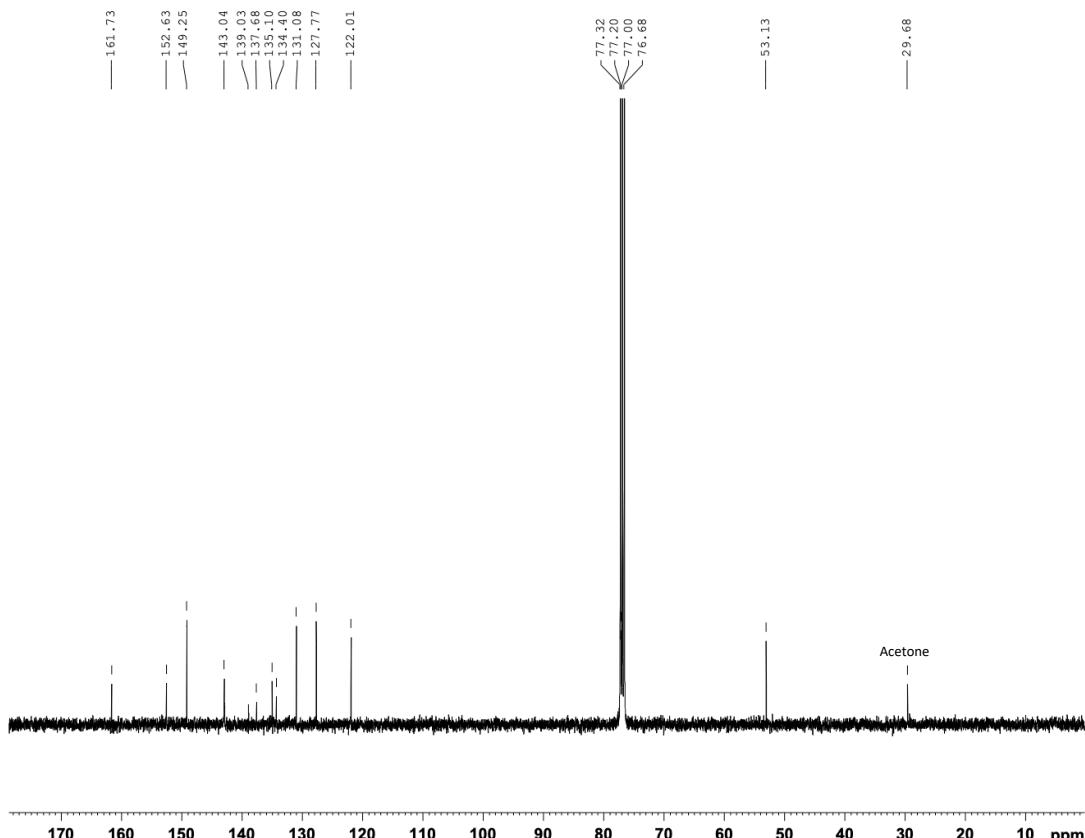


### Methyl 3-(pyridin-4-yl)thieno[3,2-*b*]pyridine-2-carboxylate (2g)

<sup>1</sup>H NMR ( $\text{CDCl}_3$ , 400 MHz) of compound **2g**

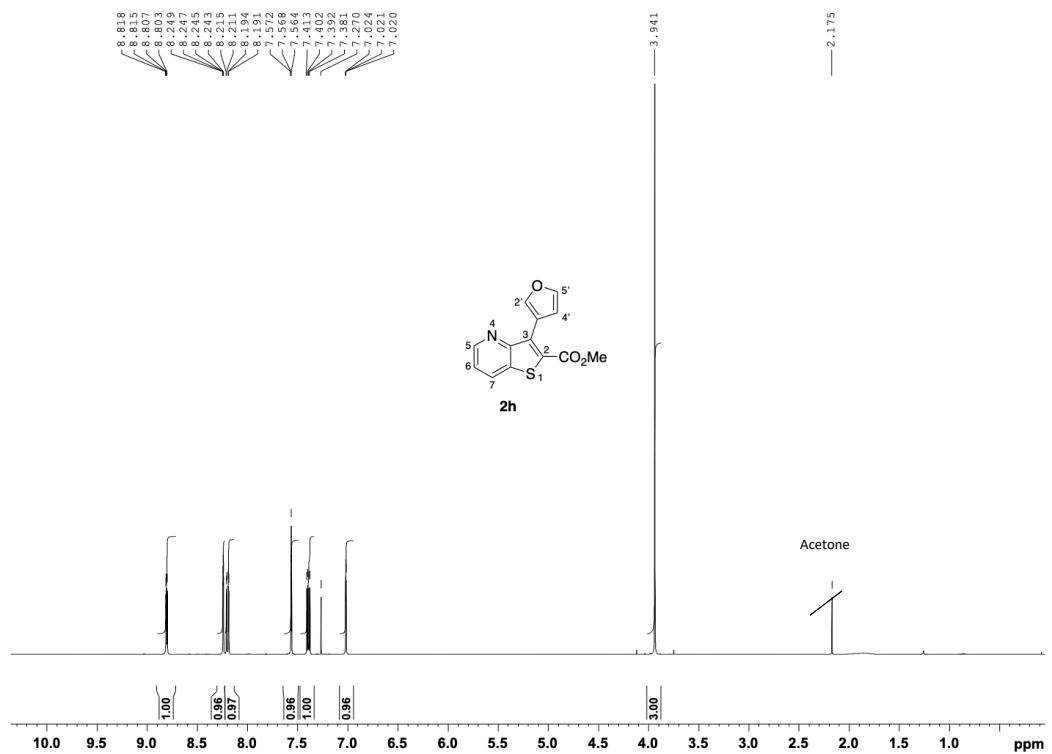


<sup>13</sup>C NMR ( $\text{CDCl}_3$ , 100.6 MHz) of compound **2g**



#### Methyl 3-(furan-3-yl)thieno[3,2-*b*]pyridine-2-carboxylate (2h)

<sup>1</sup>H NMR ( $\text{CDCl}_3$ , 400 MHz) of compound **2h**



<sup>13</sup>C NMR ( $\text{CDCl}_3$ , 100.6 MHz) of compound **2h**

