

Supporting Information

Substituted 4-oxo-crotonic acid derivatives as a new class of Protein kinase B (PknB) inhibitors: synthesis and SAR study

Changliang Xu,^{a,b,†} Xiaoguang Bai,^{a,†} Jian Xu,^a Jinfeng Ren,^a Yun Xing,^a Ziqiang Li,^a Juxian Wang,^a Jingjing Shi,^b Liyan Yu^{*,a} and Yucheng Wang^{*,a}

^aInstitute of Medicinal Biotechnology, Chinese Academy of Medical Science & Peking Union Medical College, Beijing 100050, China. E-mail address: yly@cpcc.ac.cn (L.Y.Y.); wyc9999@gmail.com (Y.C.W.); Tel./Fax: +86-010-6318-7118 (L.Y.Y.); +86-010-6316-5263 (Y.C.W.).

^bJiangsu Protein Drug Engineering Laboratory, Food & Drug Analysis and Testing Center, Jiangsu Food & Pharmaceutical Science College, Huai'an 223003, Jiangsu, China. E-mail address: clxu1986@163.com; Tel: +86-0517-87088209.

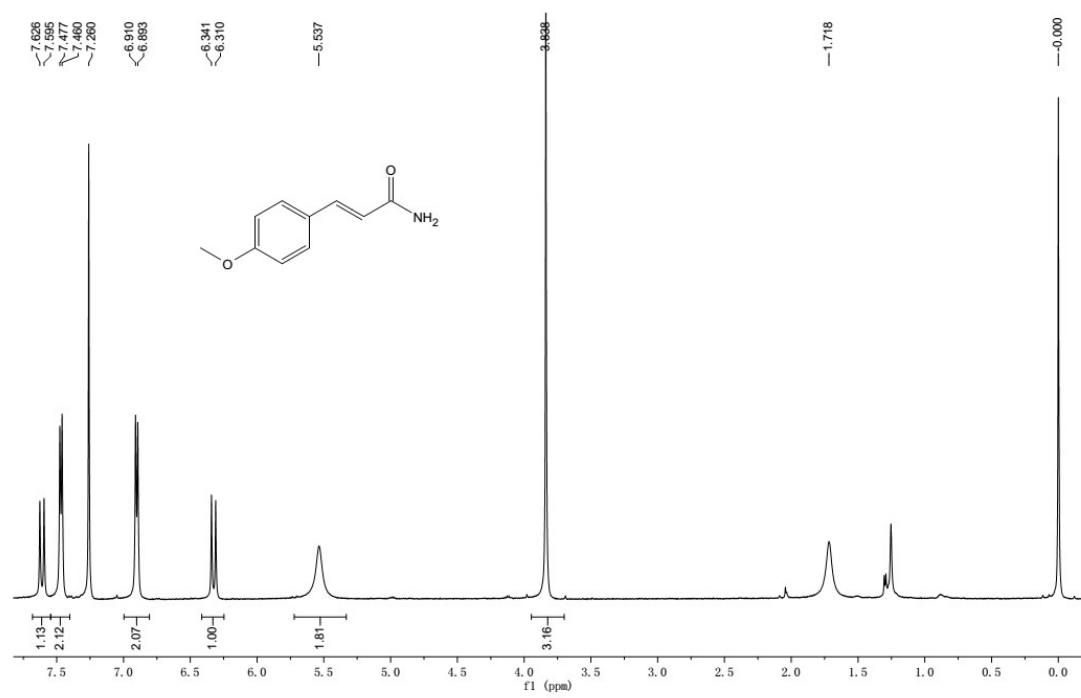
† These authors contributed equally to this work.

NMR spectra of synthesized compounds:

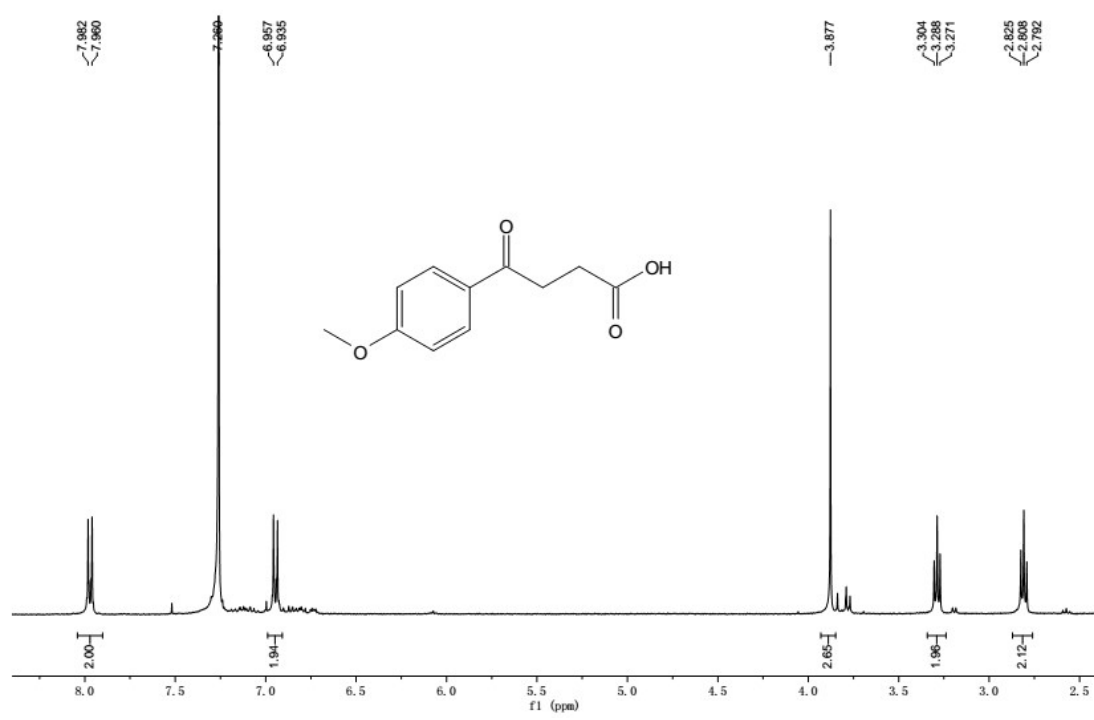
1b



1c



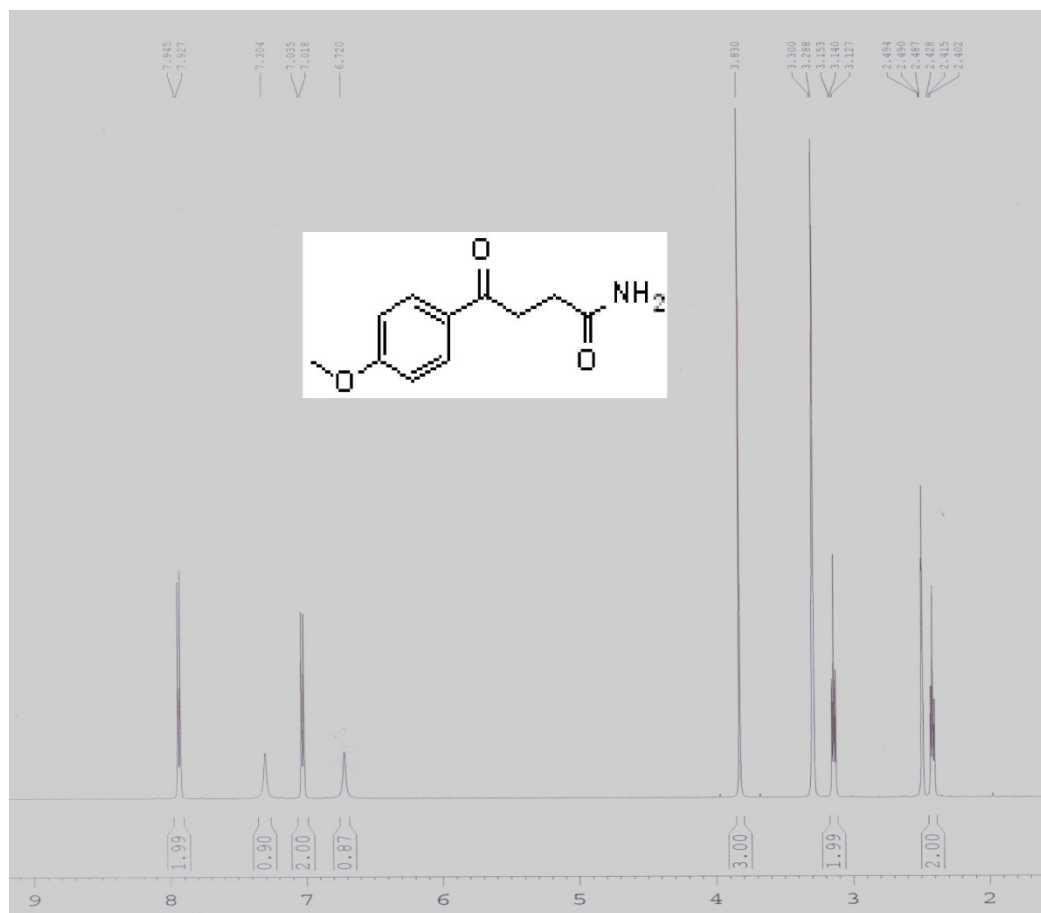
2a



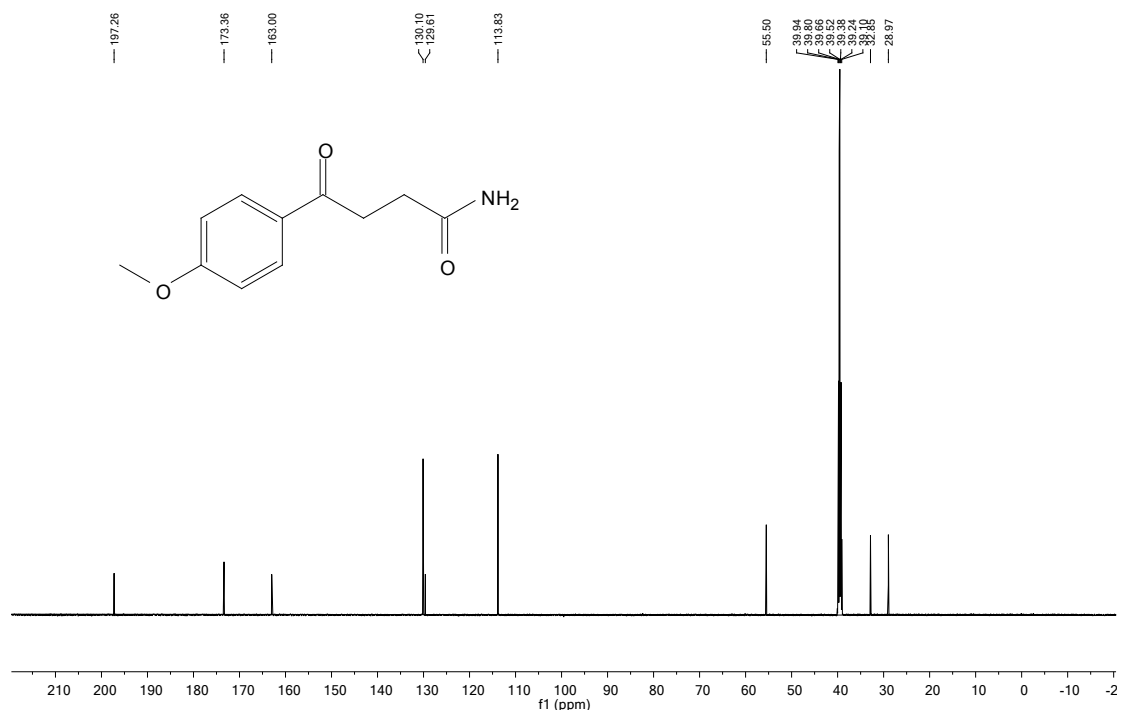
2b



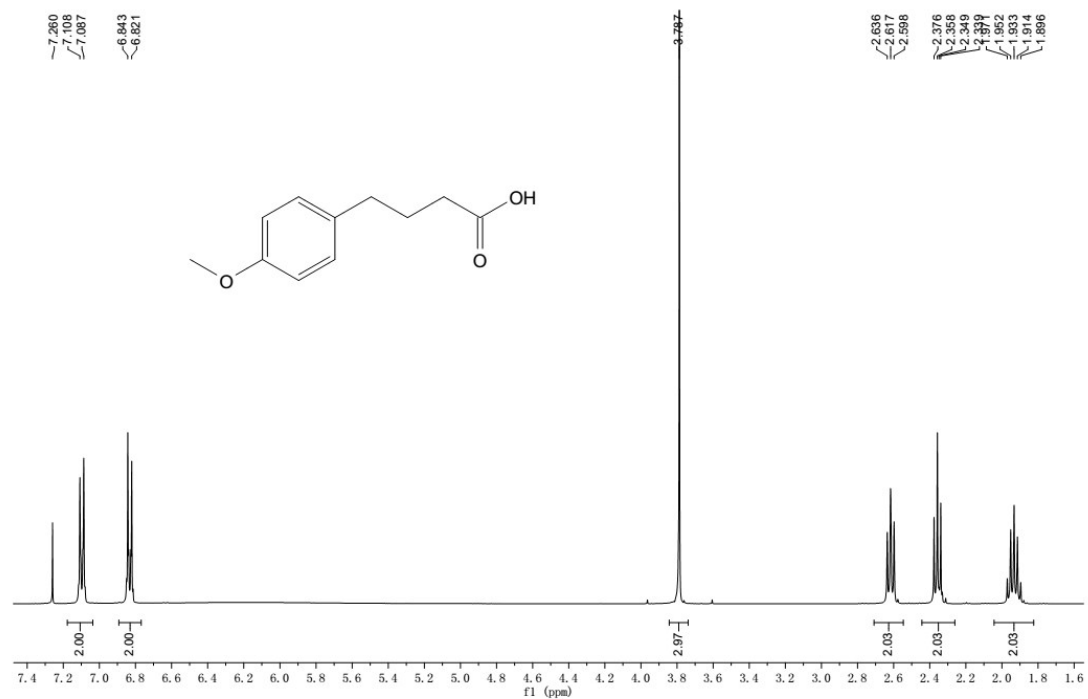
2c



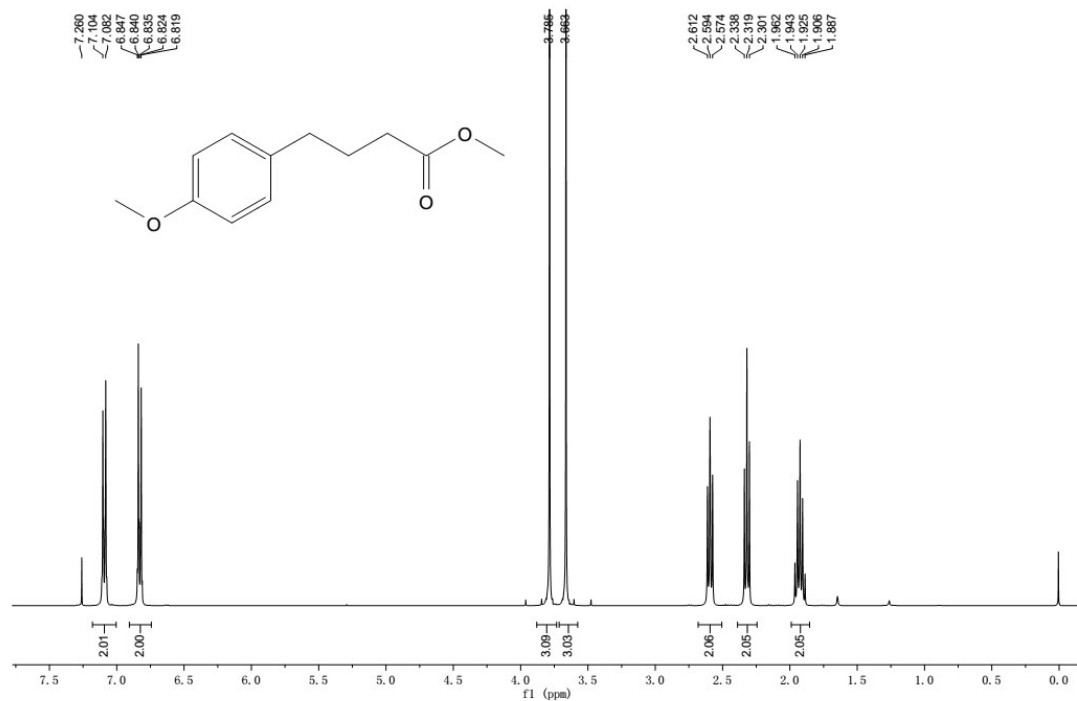
BRUKER AVANCE III 13C-NMR 2c IN DMSO-d6



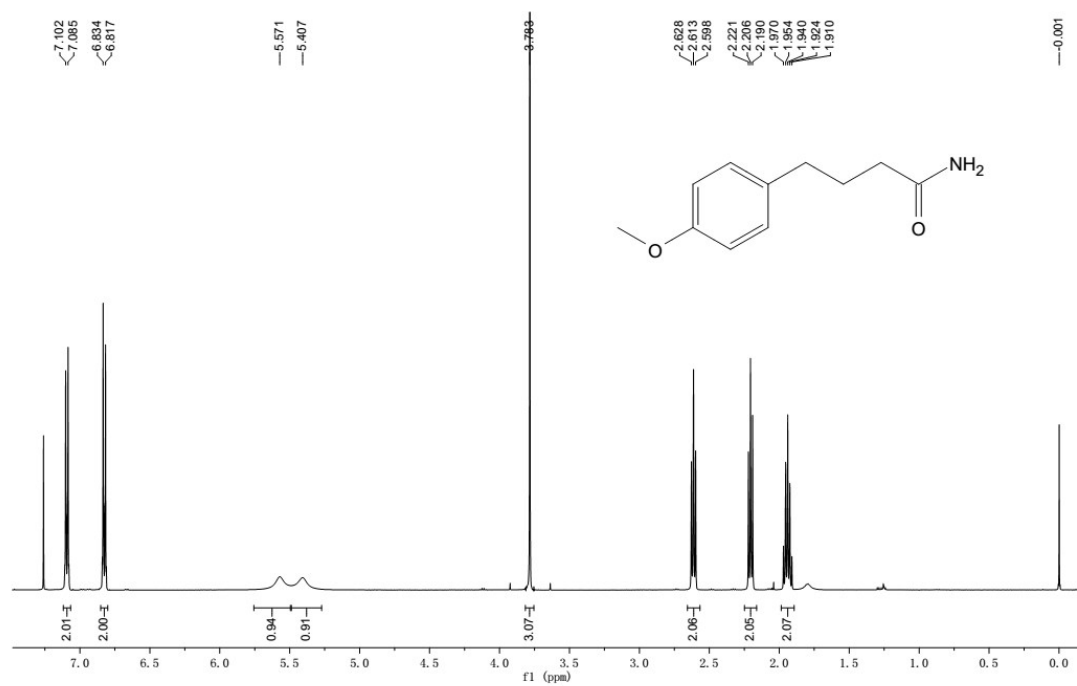
3a



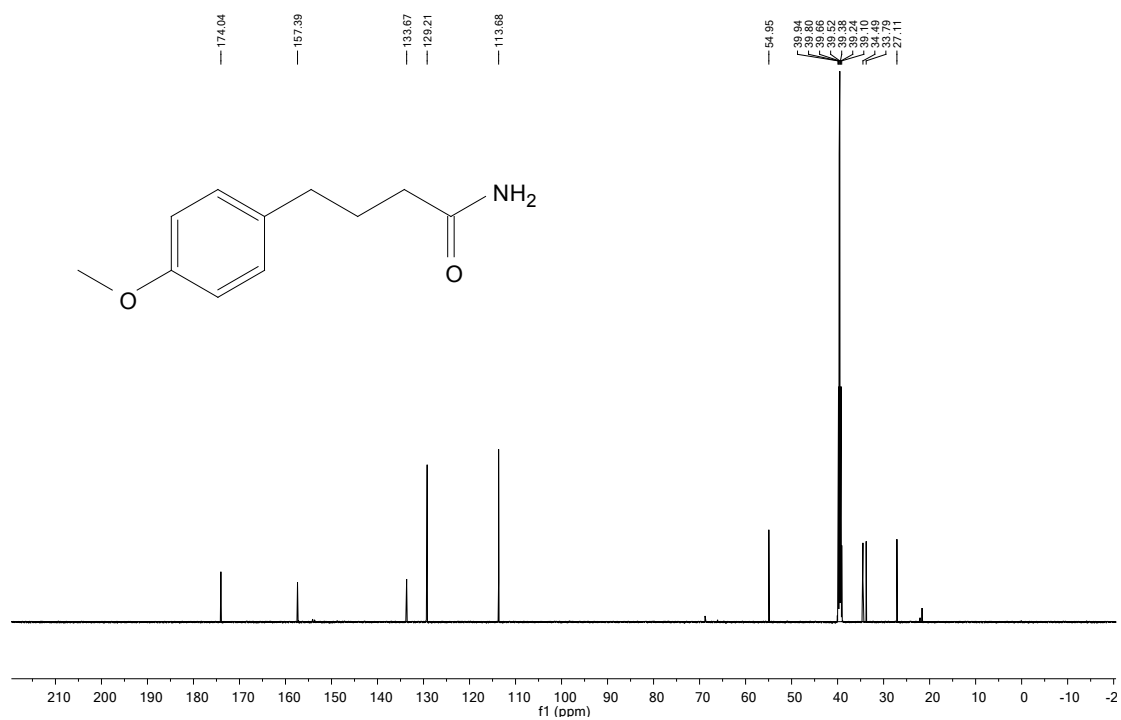
3b



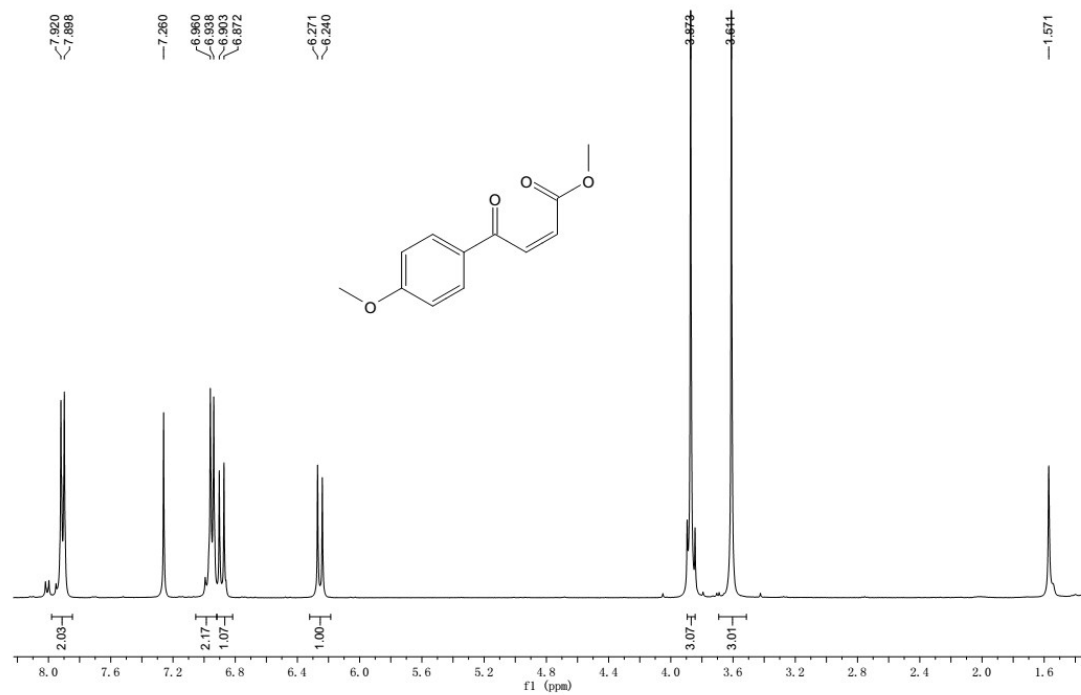
3c



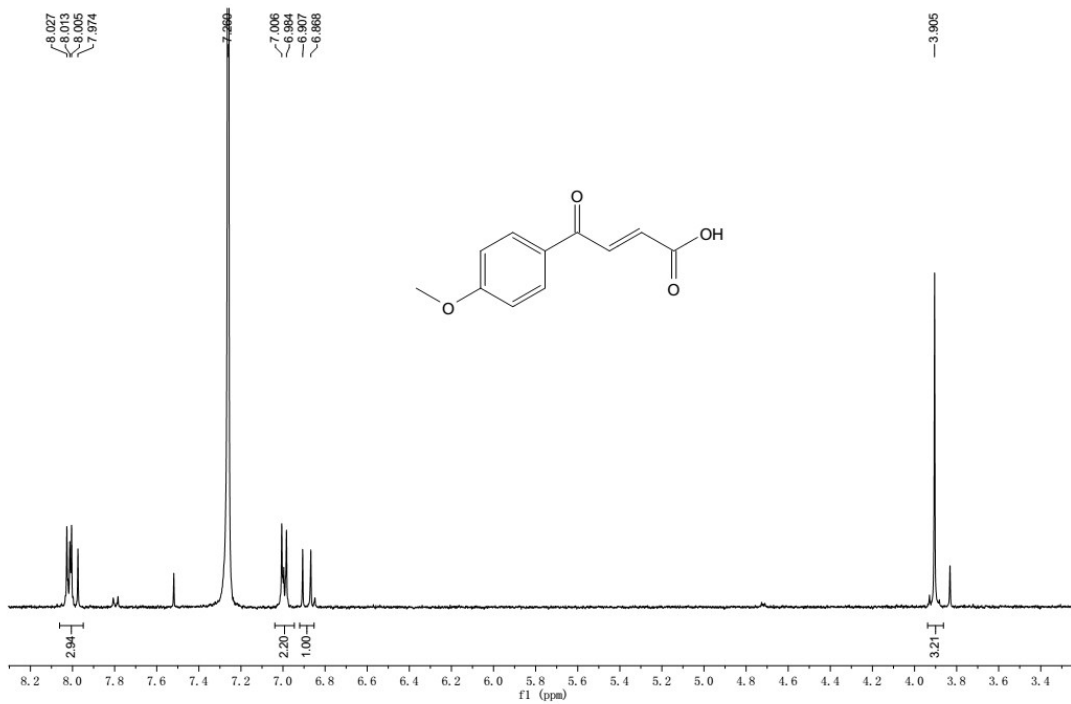
BRUKER AVANCE III 13C-NMR 3c IN DMSO-d6



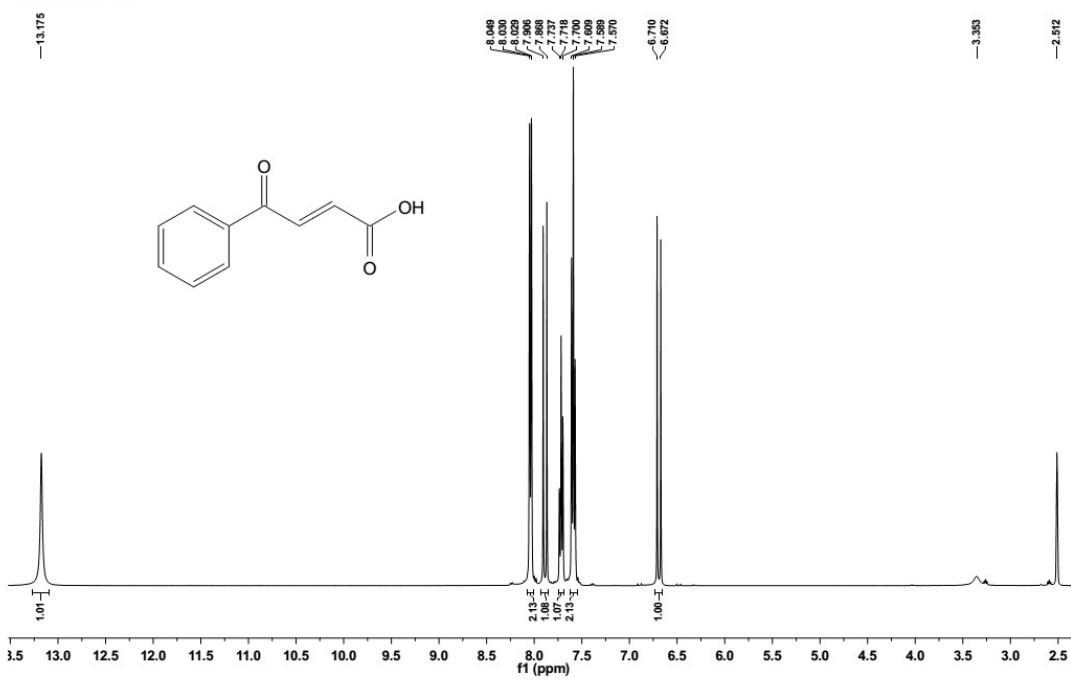
4



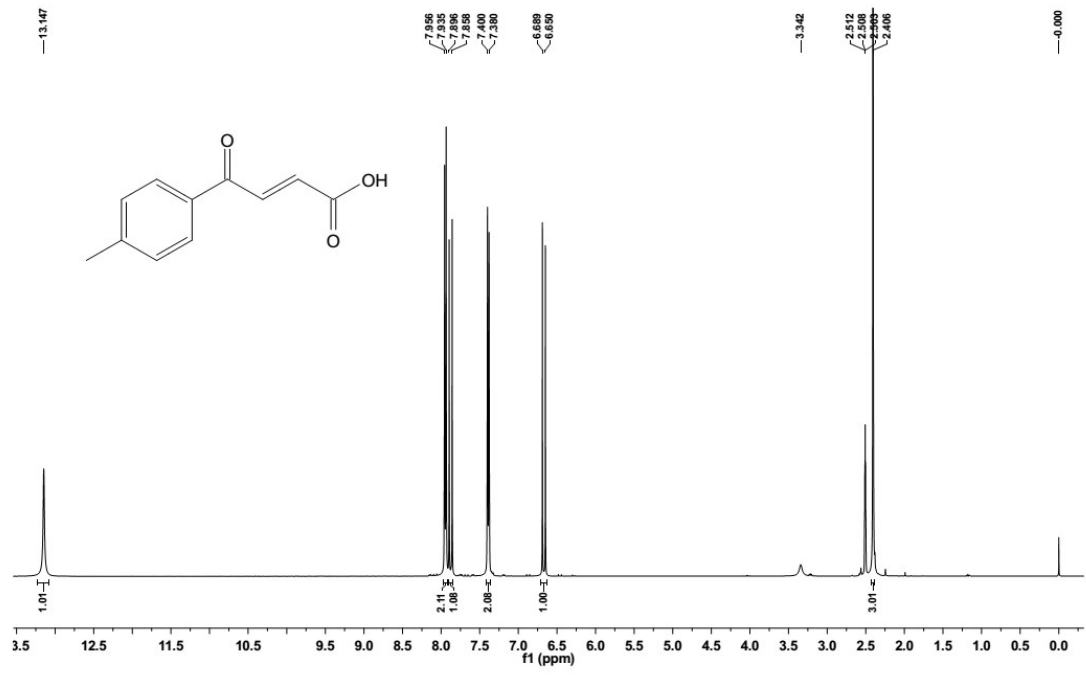
7a



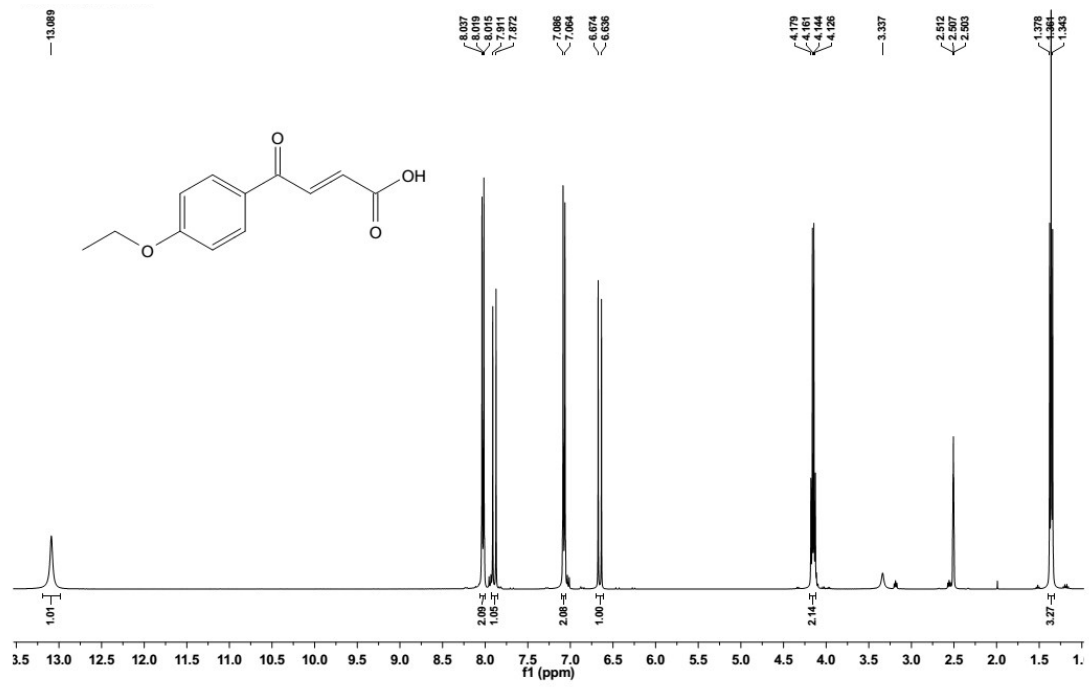
7b



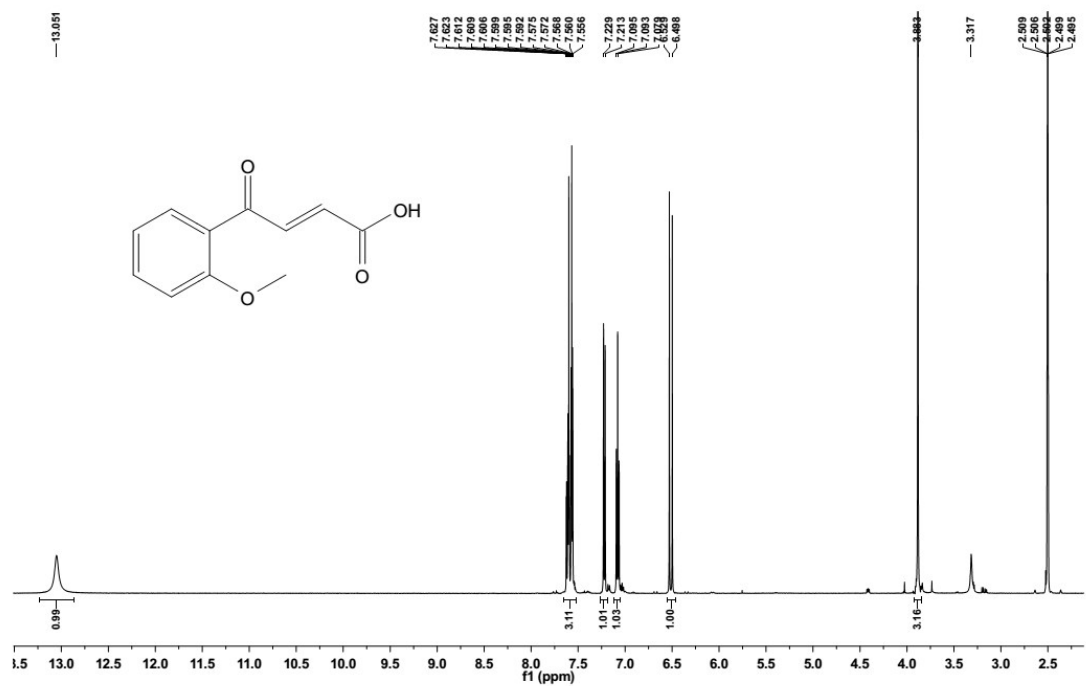
7c



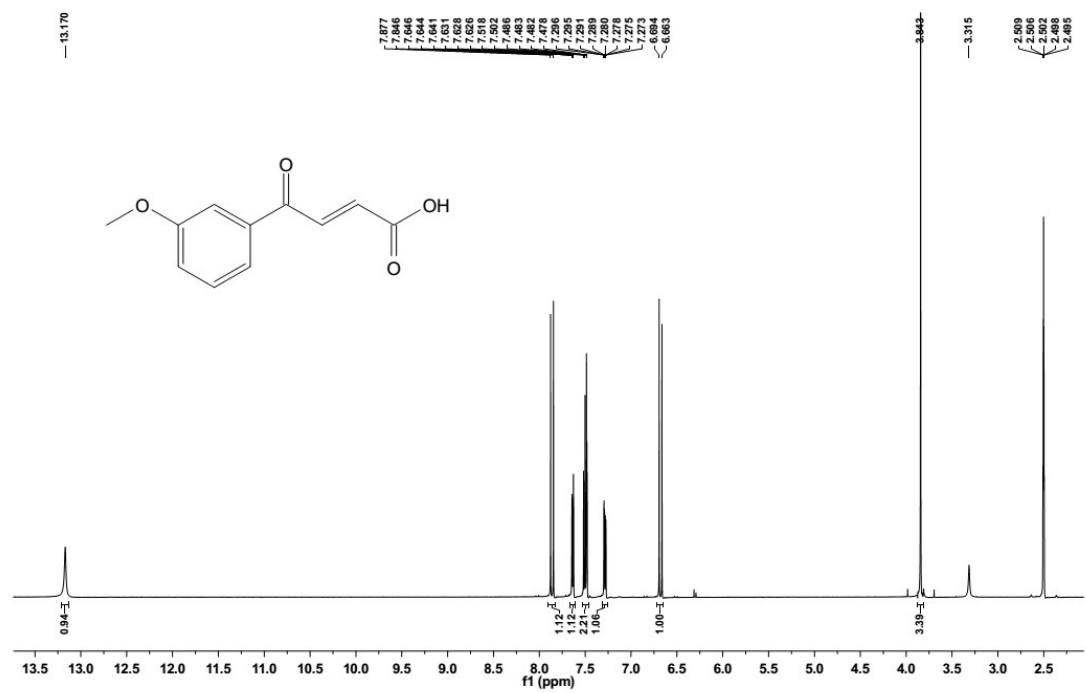
7d



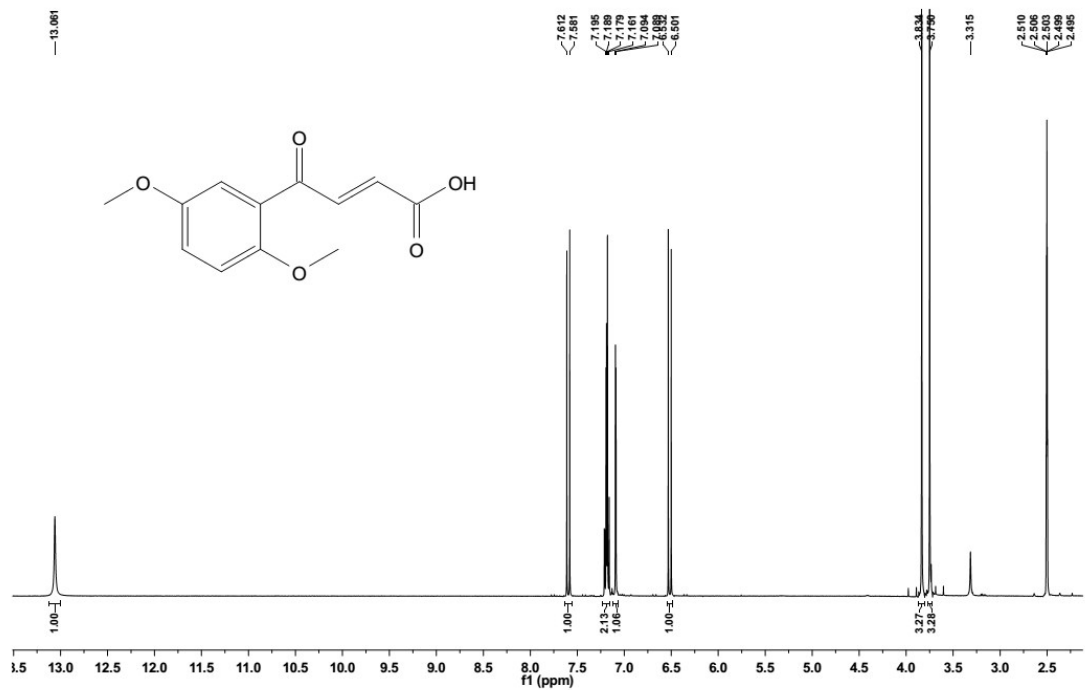
7e



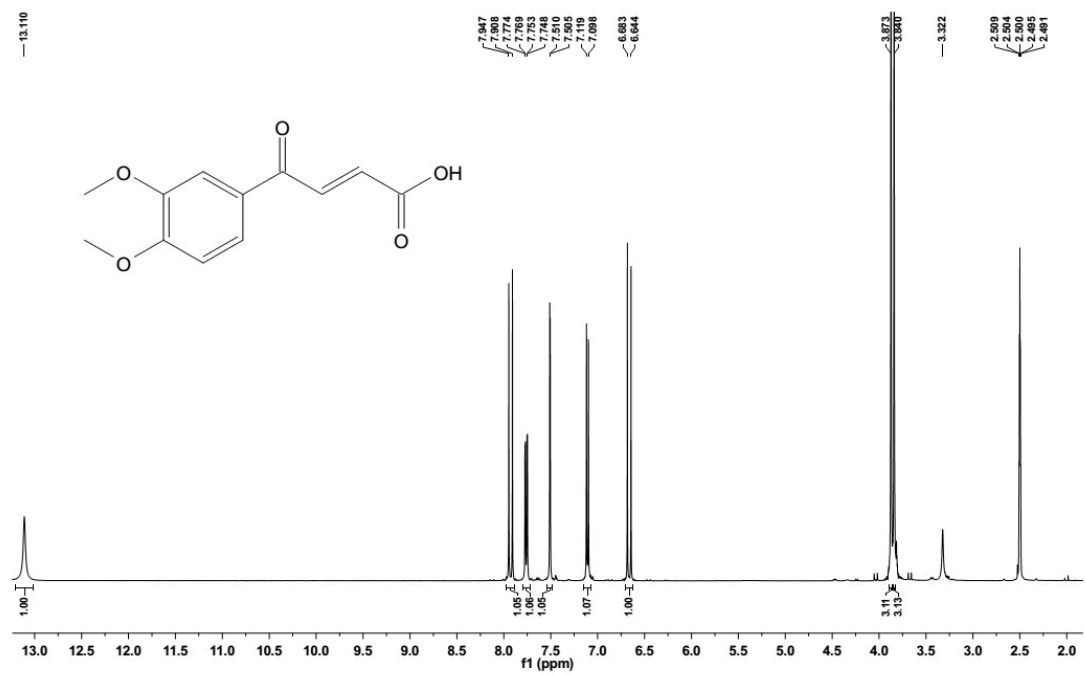
7f



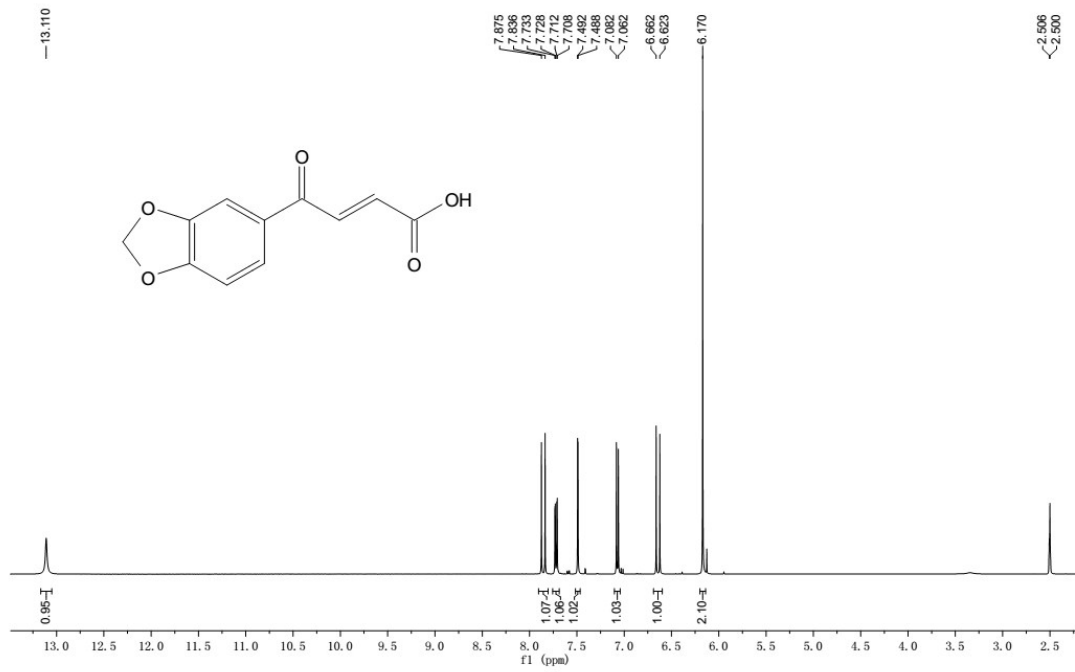
7g



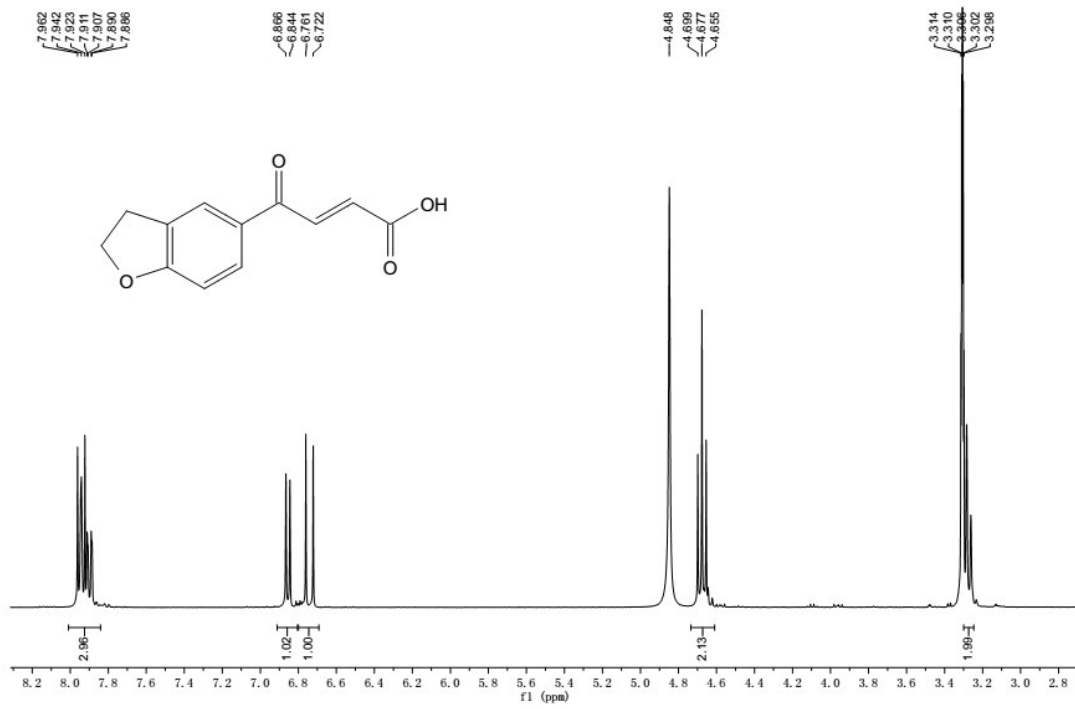
7h



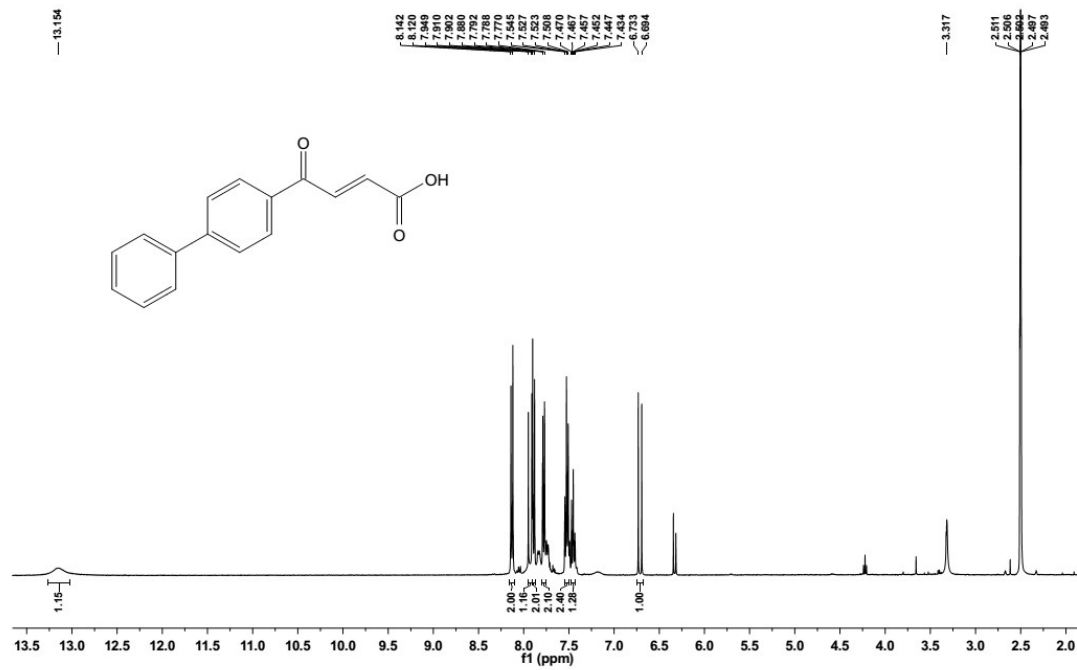
7i



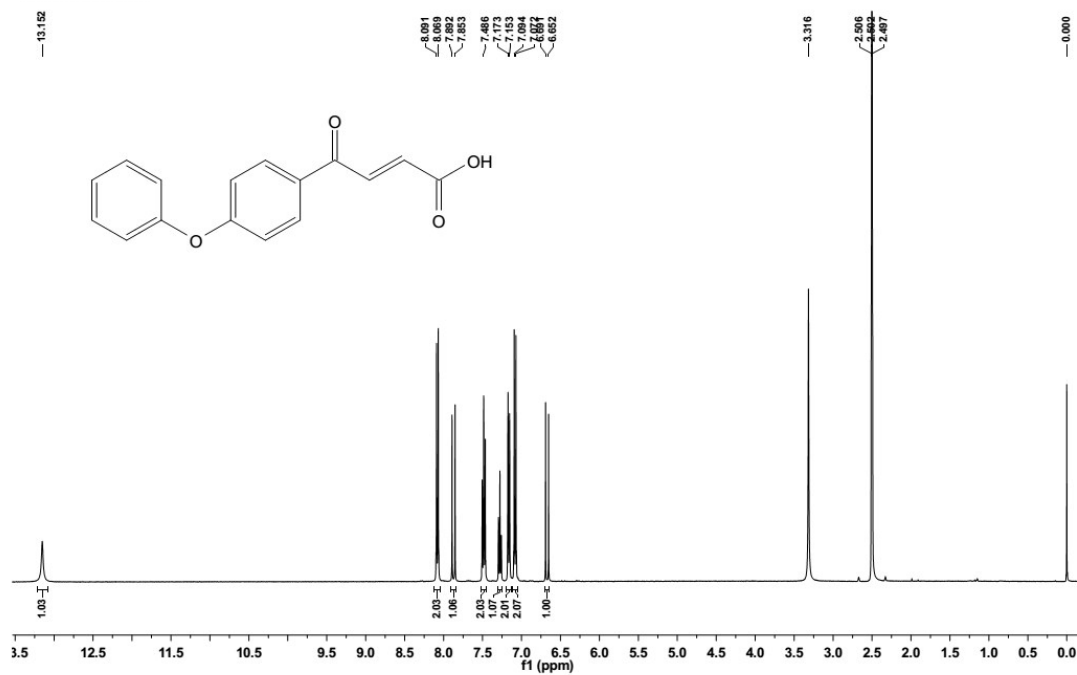
7j



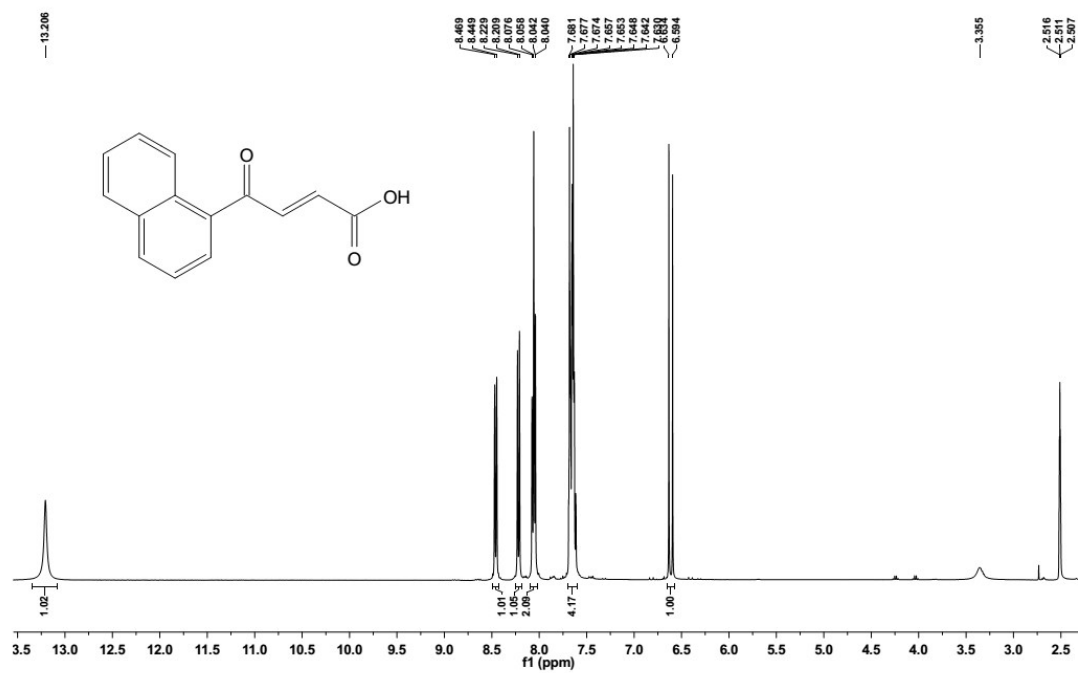
7k



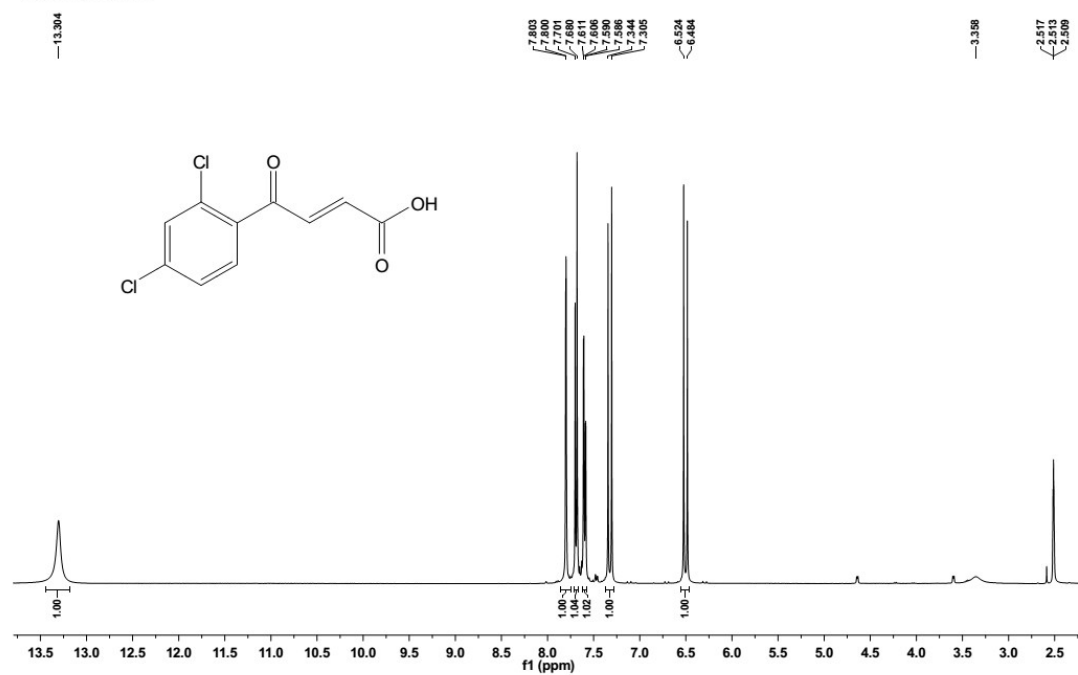
7l



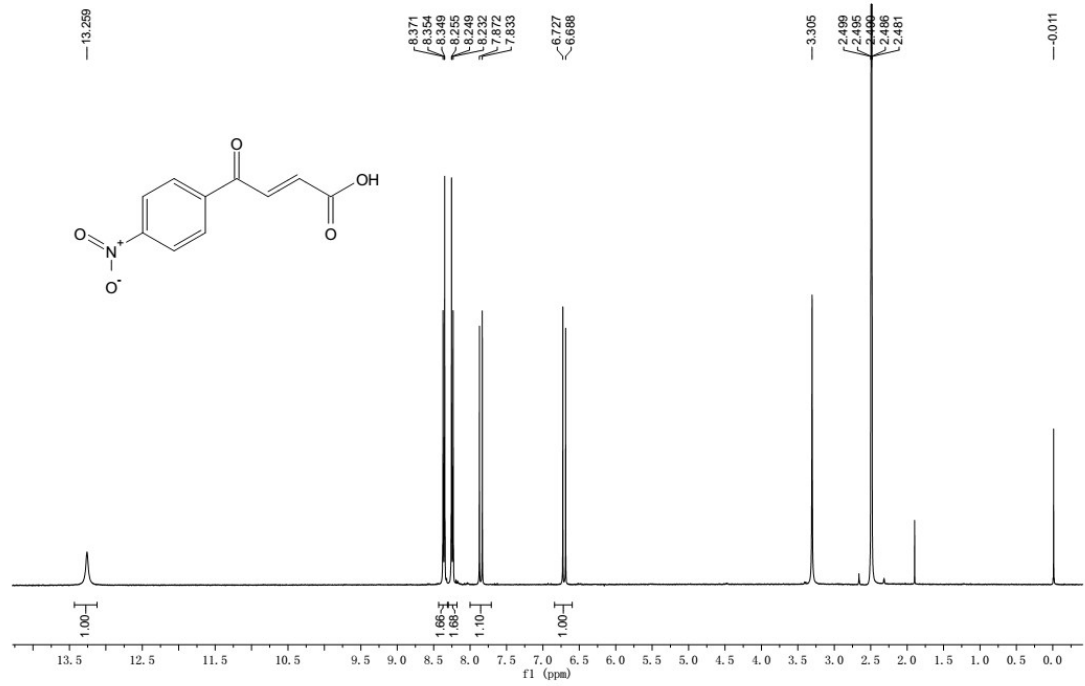
7m



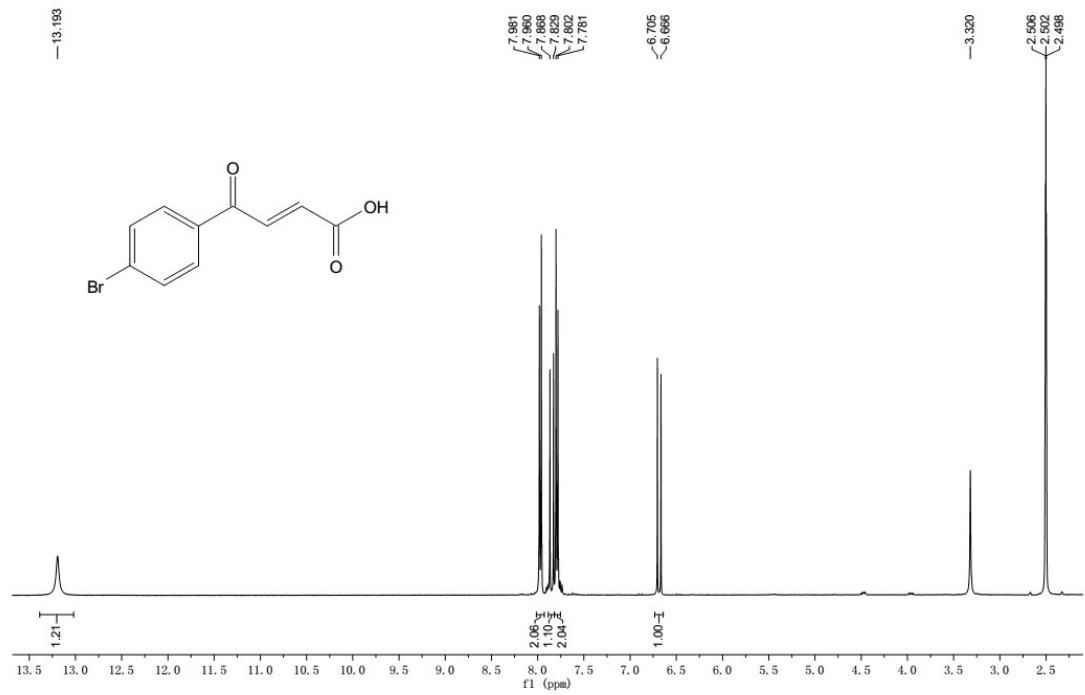
7n



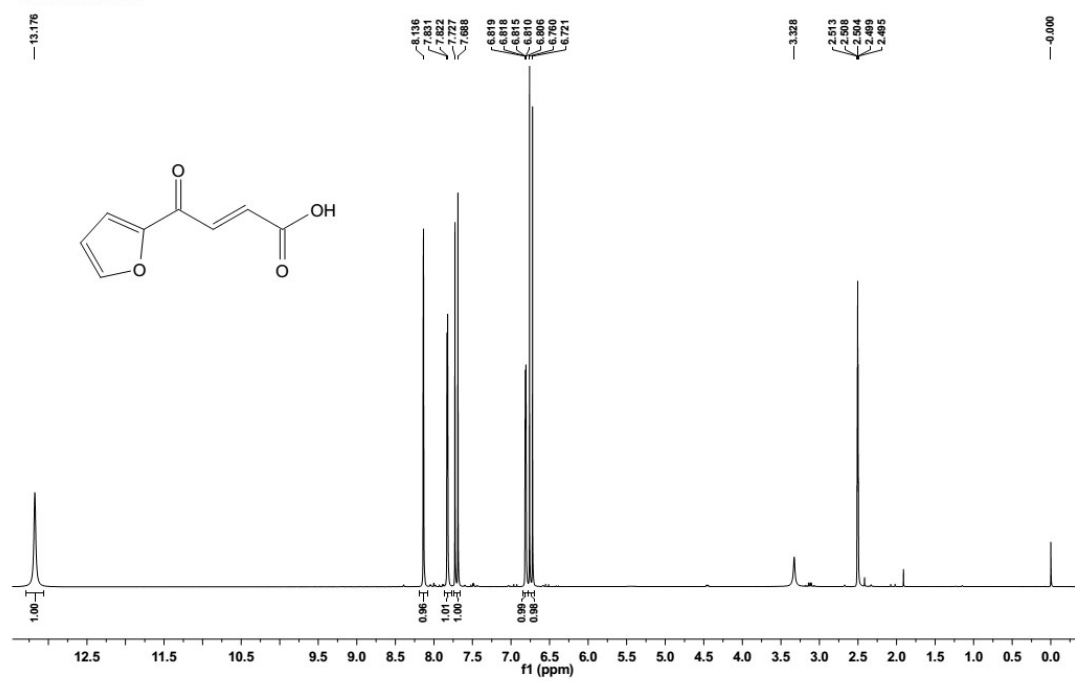
7o



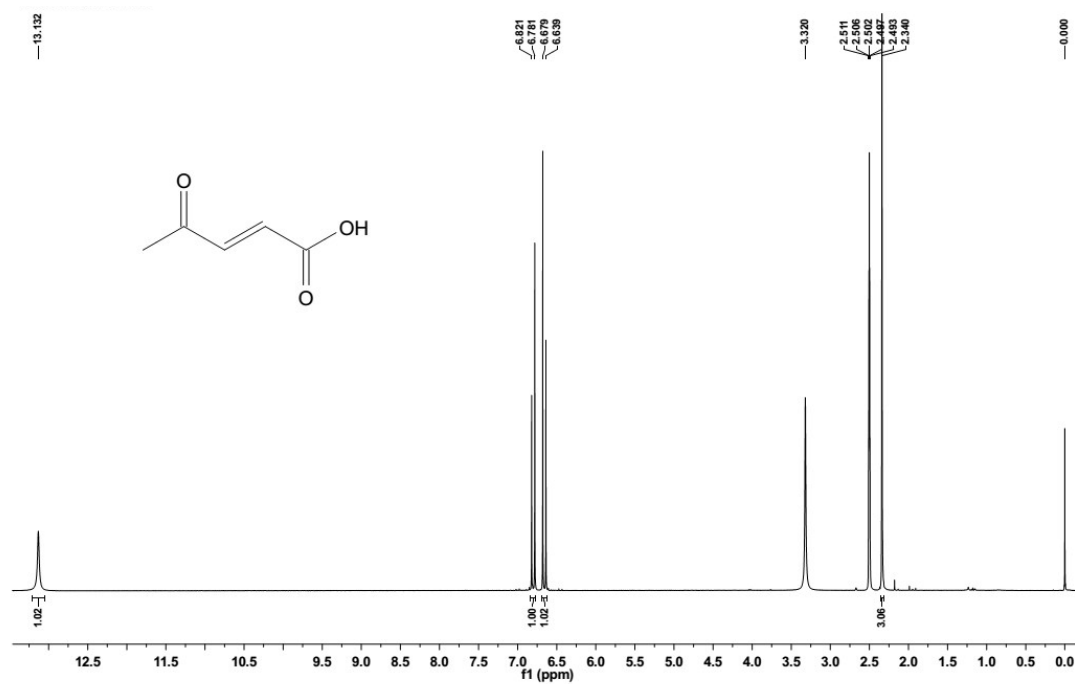
7p



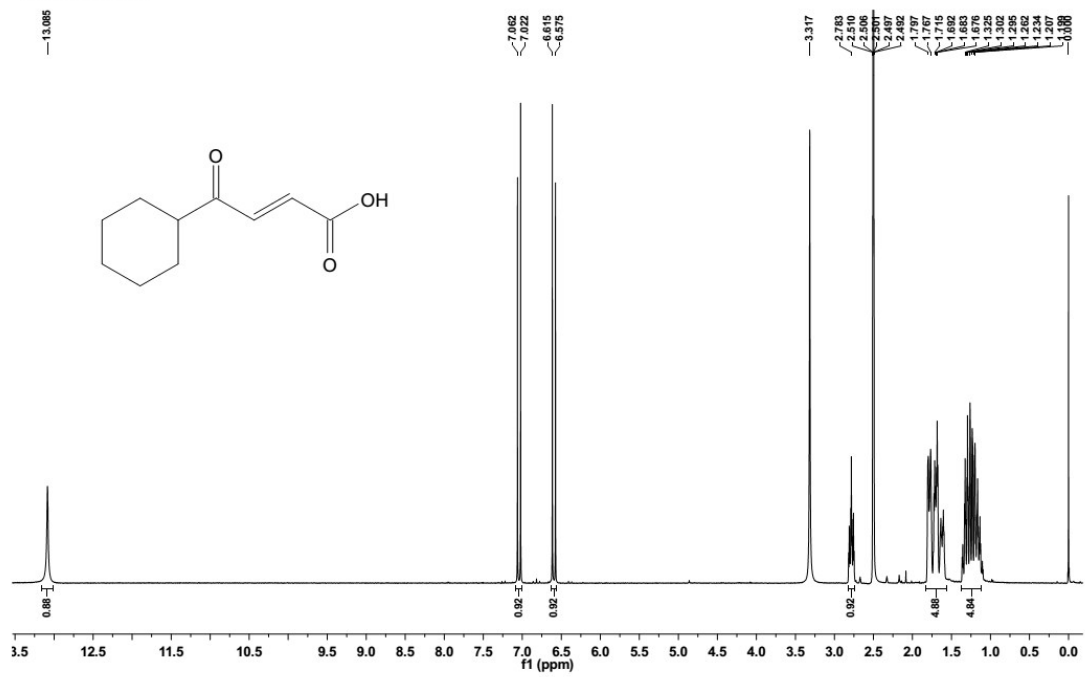
7q



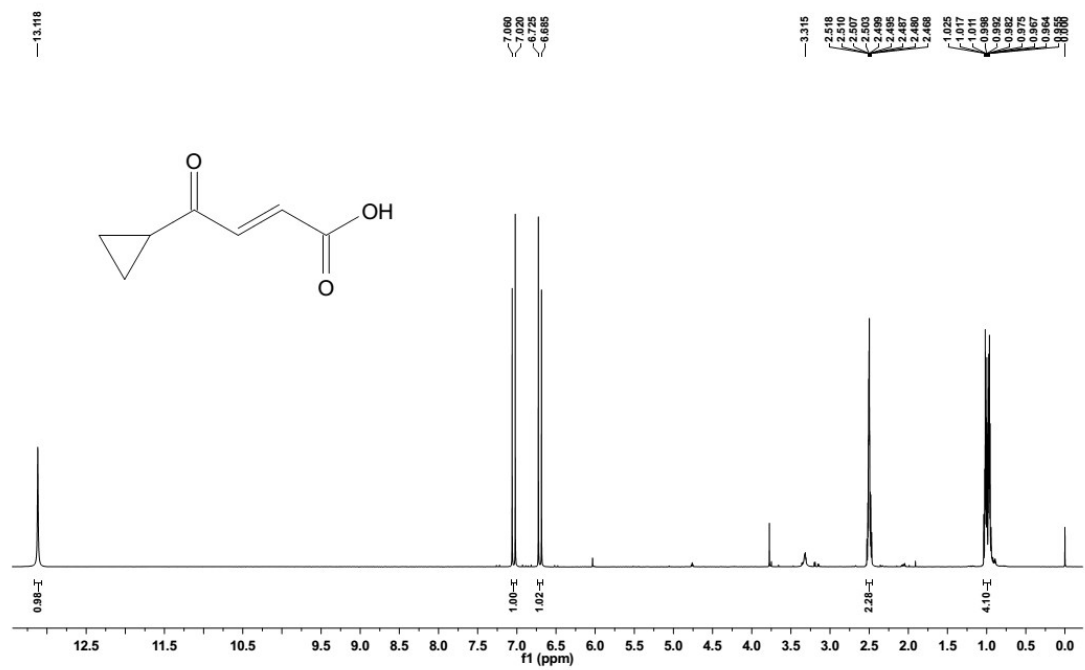
7r



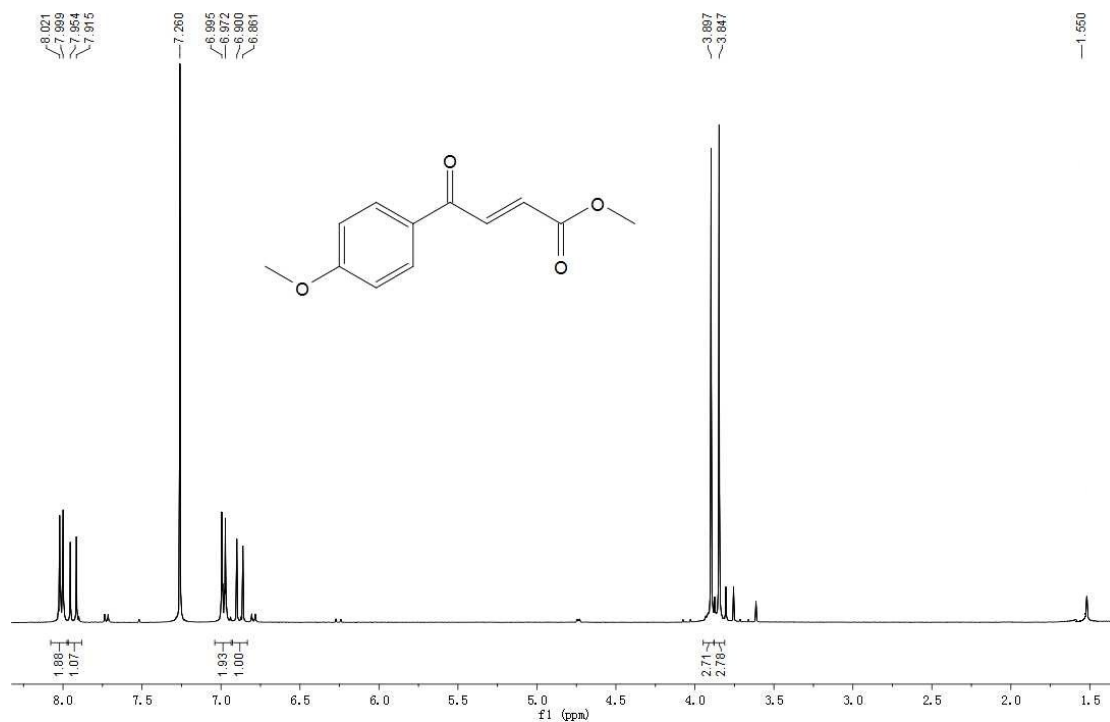
7s



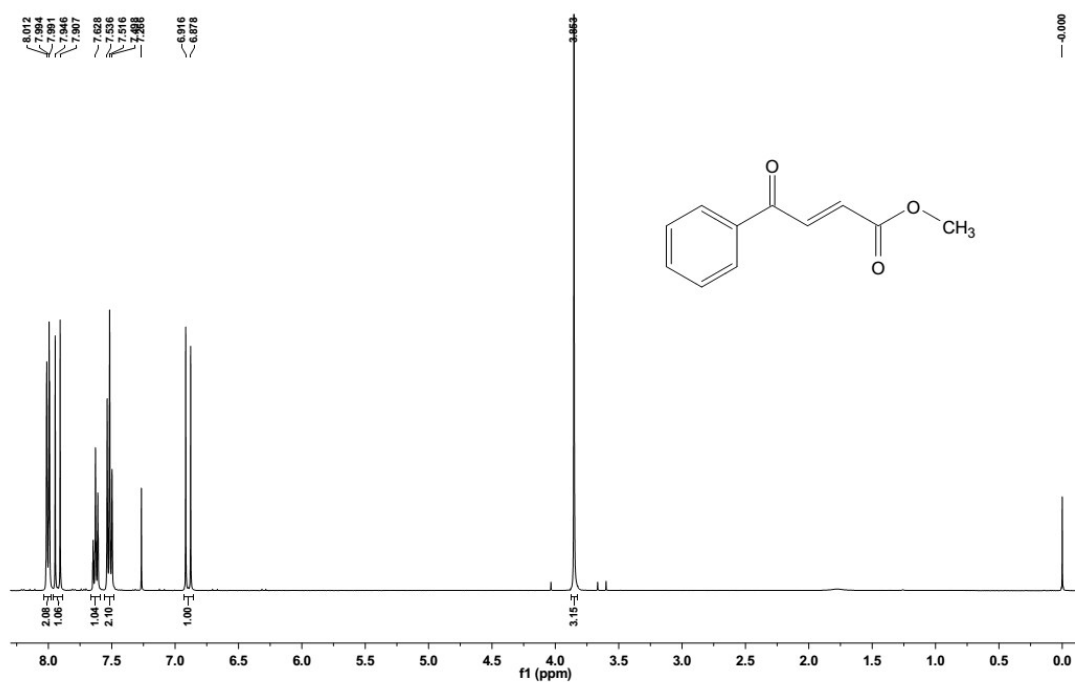
7t



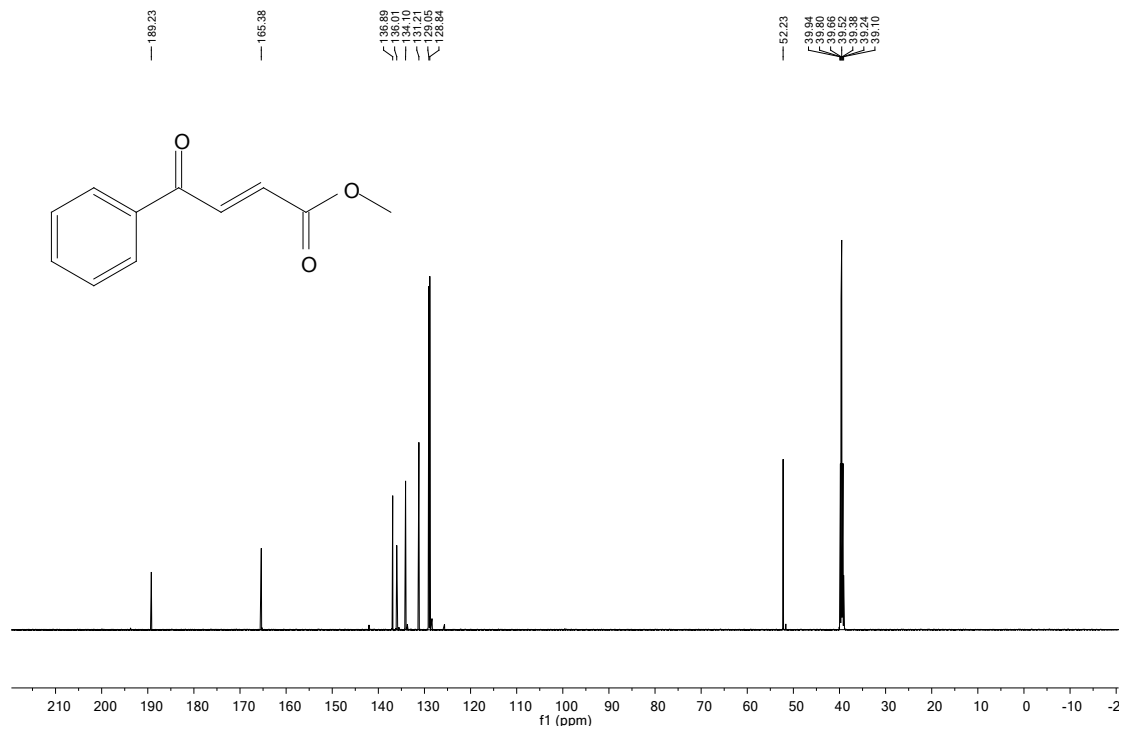
8a



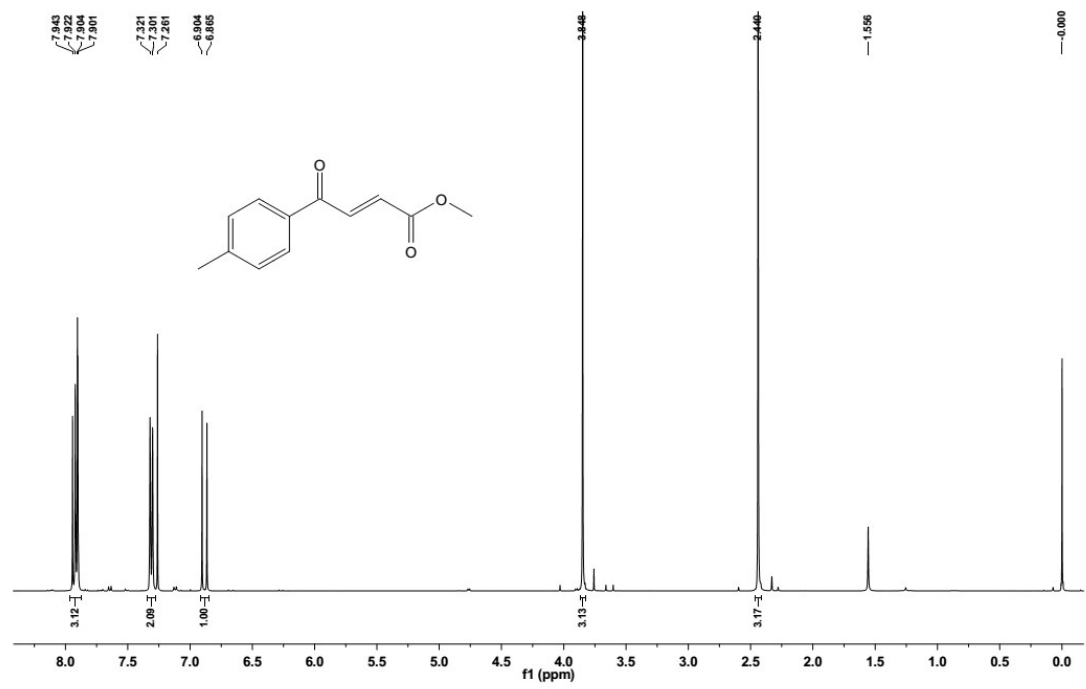
8b



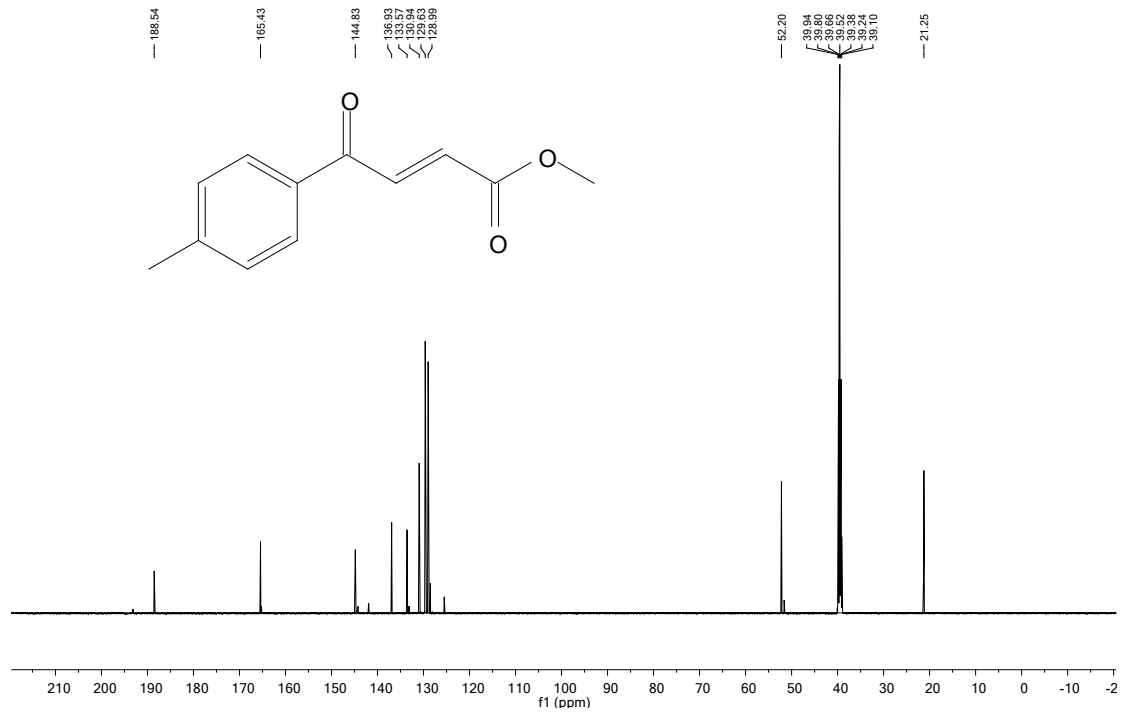
BRUKER AVANCE III 13C-NMR 8b IN DMSO-d6



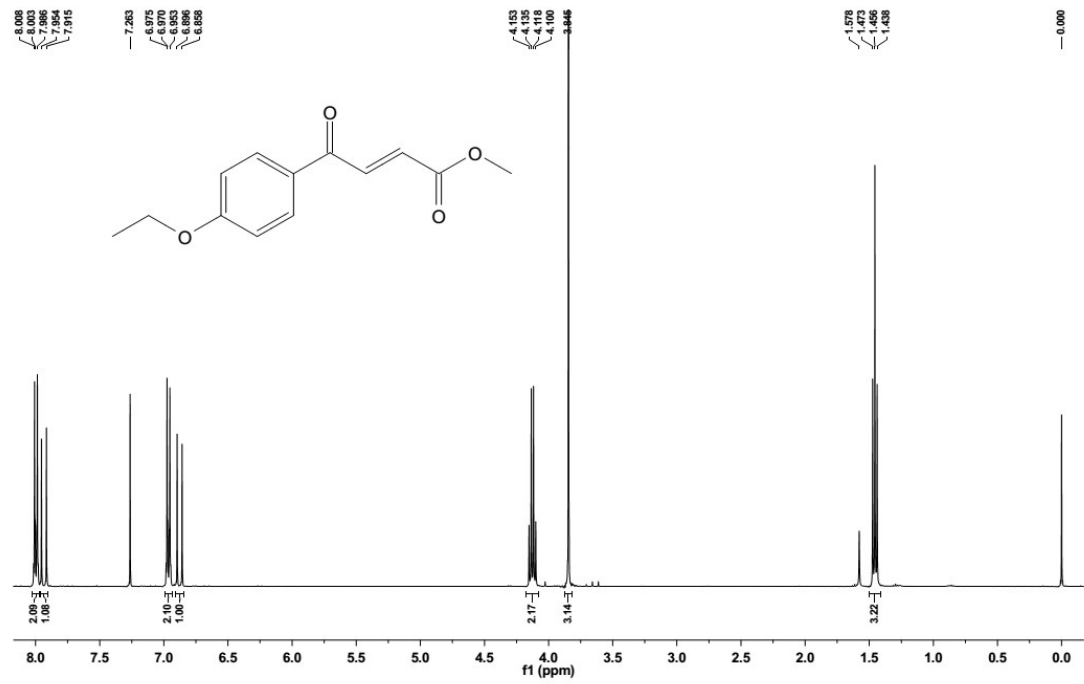
8c



BRUKER AVANCE III 13C-NMR 8c IN DMSO-d6



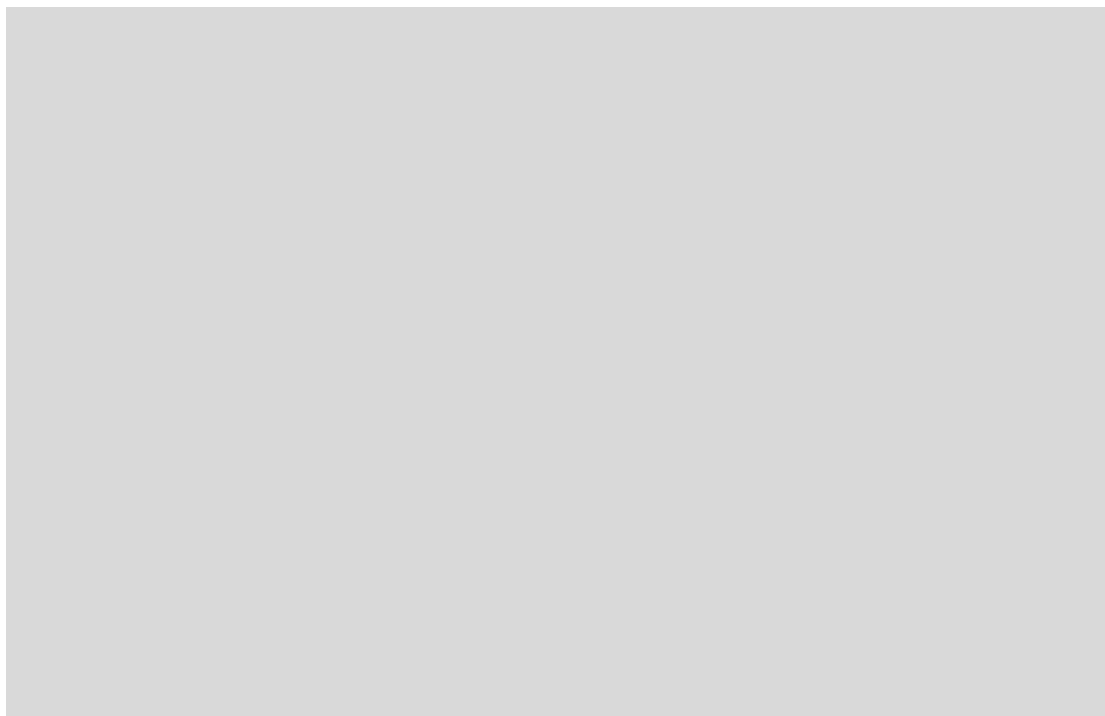
8d



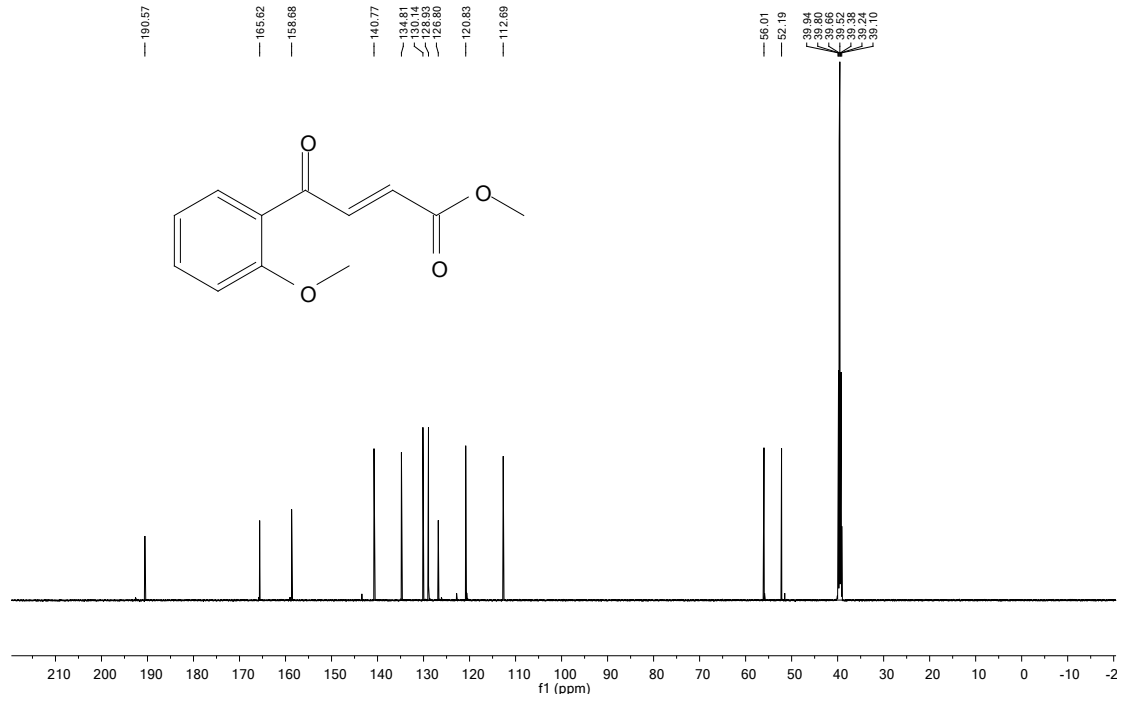
BRUKER AVANCE III 13C-NMR 8d IN DMSO-d6



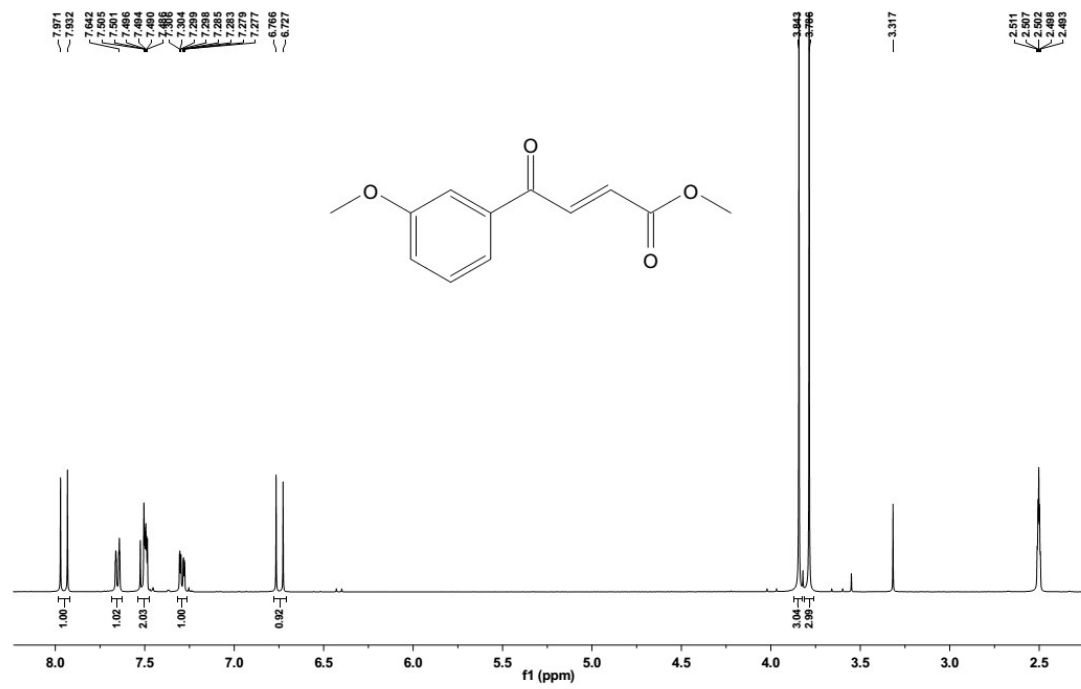
8e



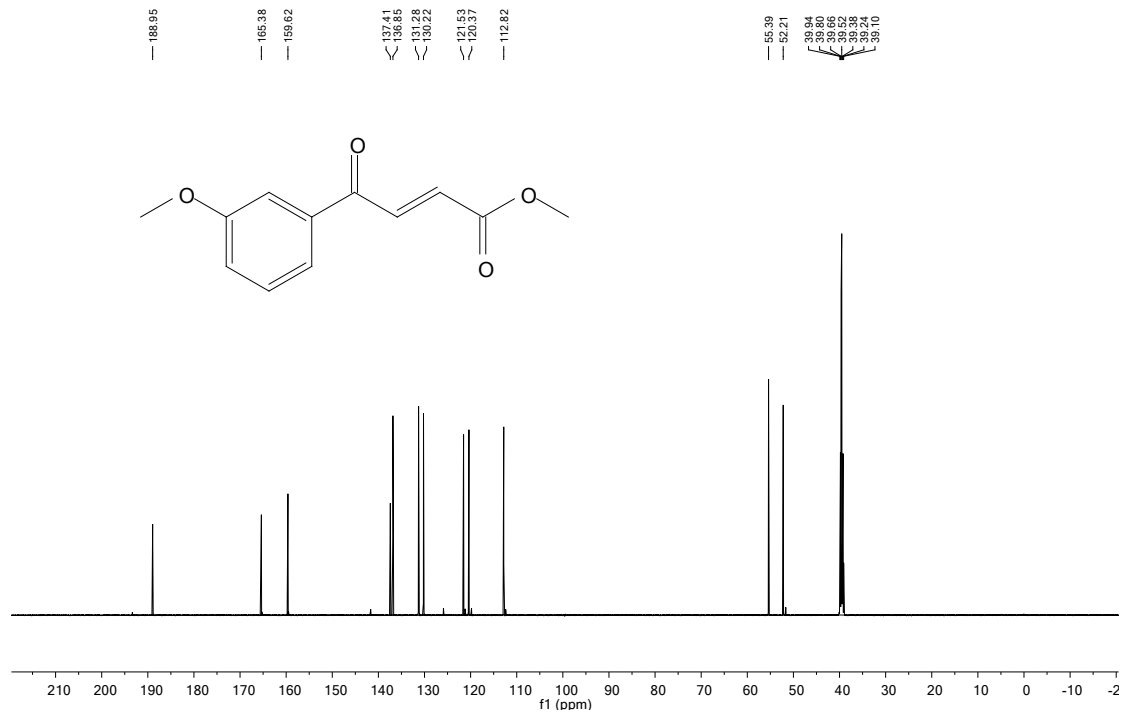
BRUKER AVANCE III 13C-NMR 8e IN DMSO-d6



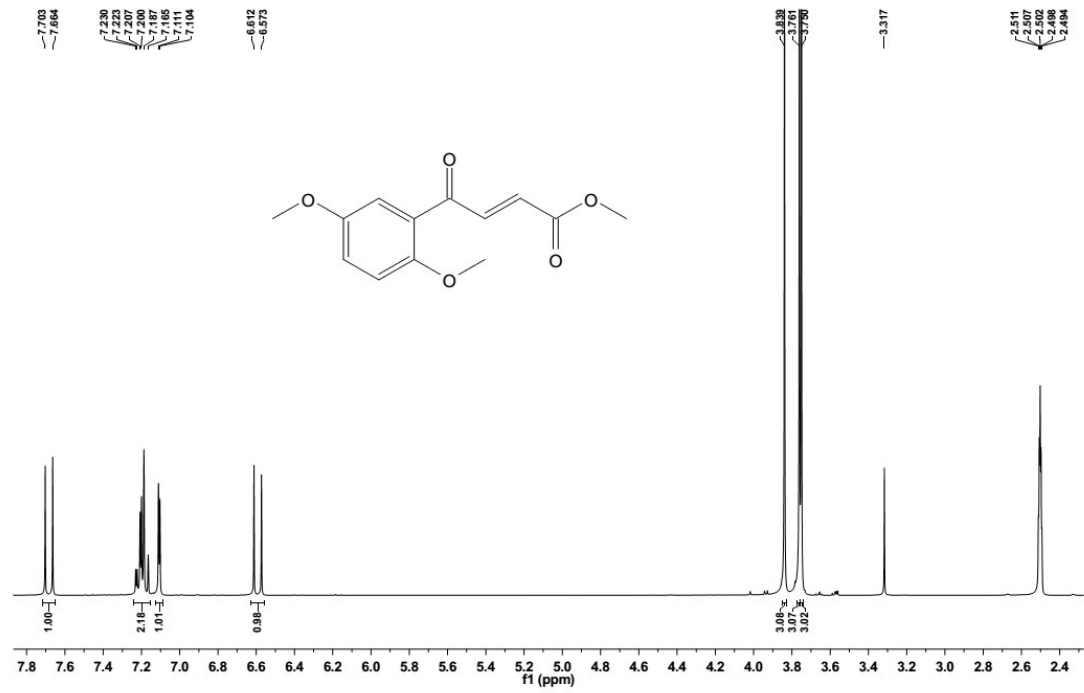
8f



BRUKER AVANCE III 13C-NMR 8f IN DMSO-d6



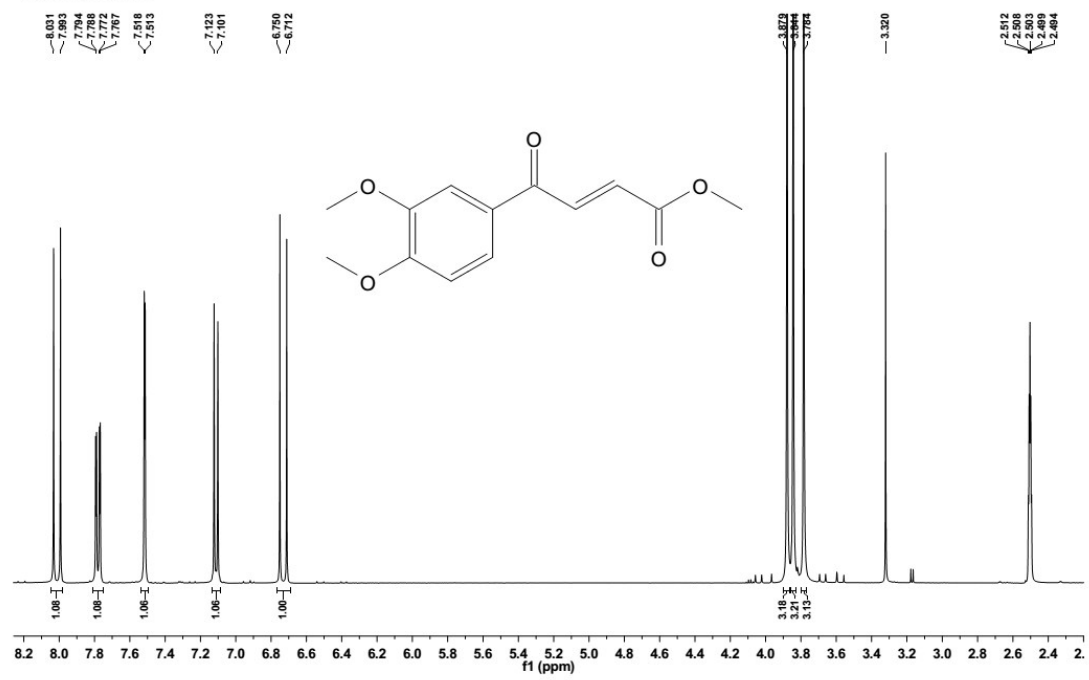
8g



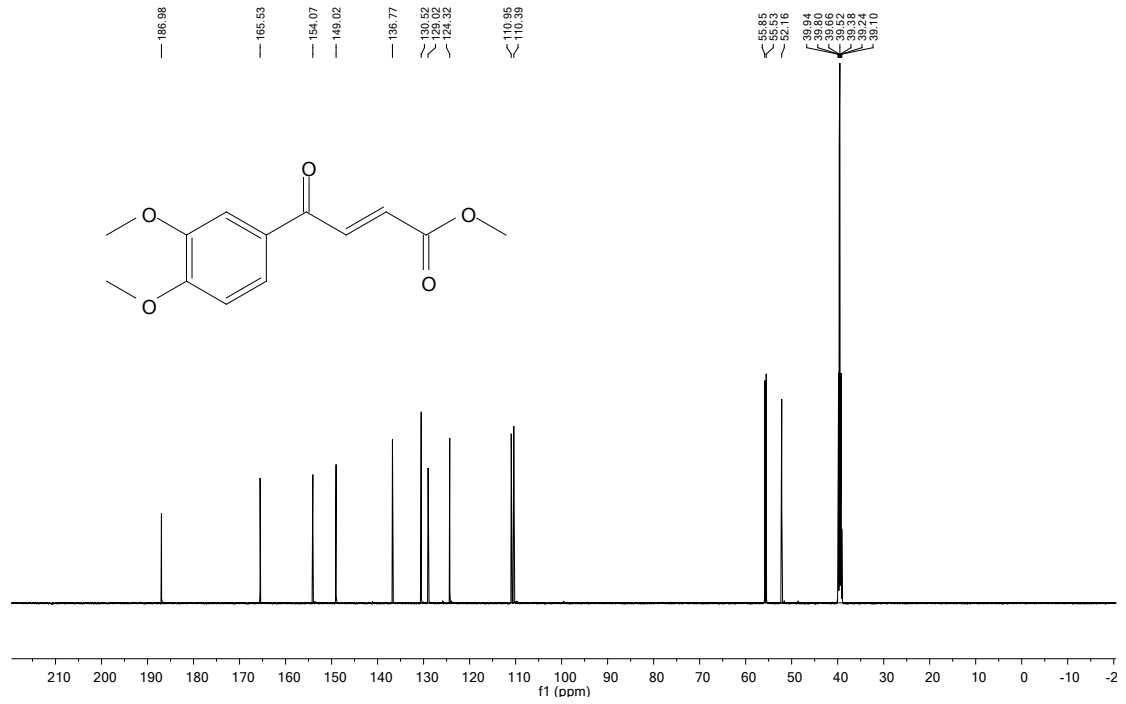
BRUKER AVANCE III 13C-NMR 8g IN DMSO-d6



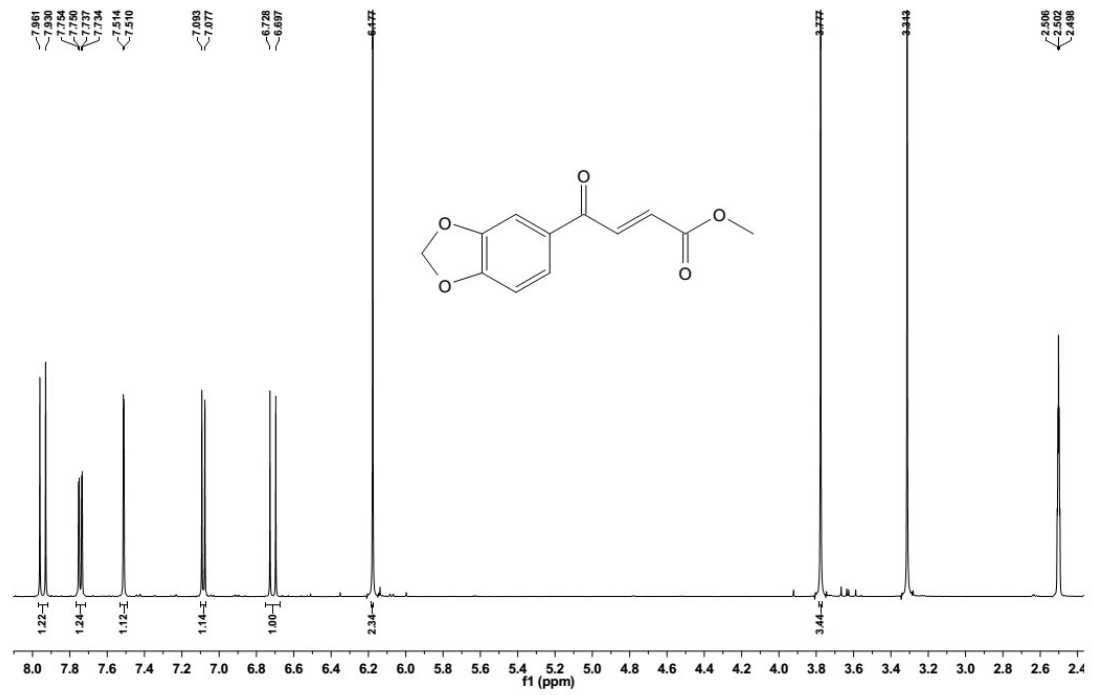
8h



BRUKER AVANCE III 13C-NMR 8h IN DMSO-d6



8i



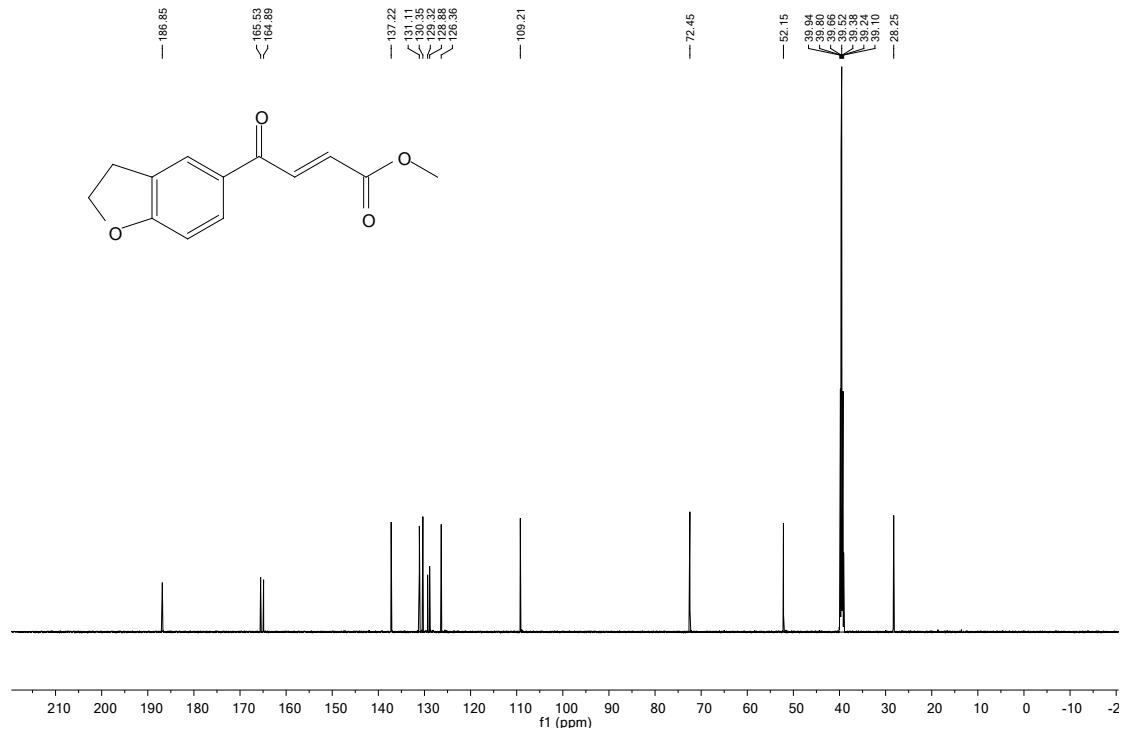
BRUKER AVANCE III 13C-NMR 8i IN DMSO-d6



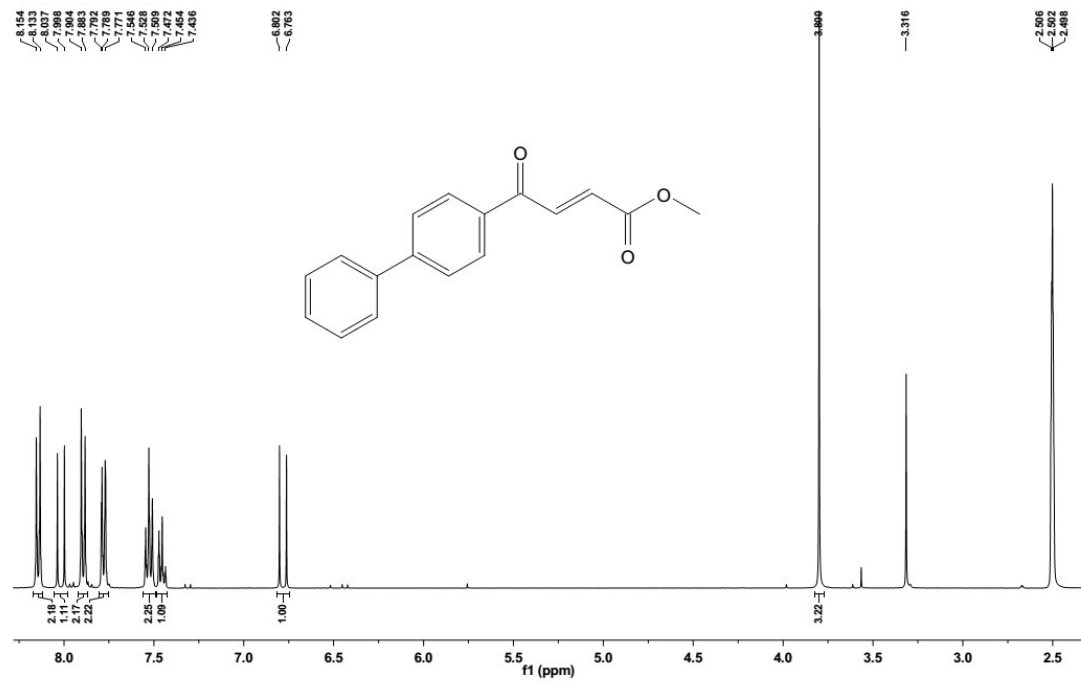
8j



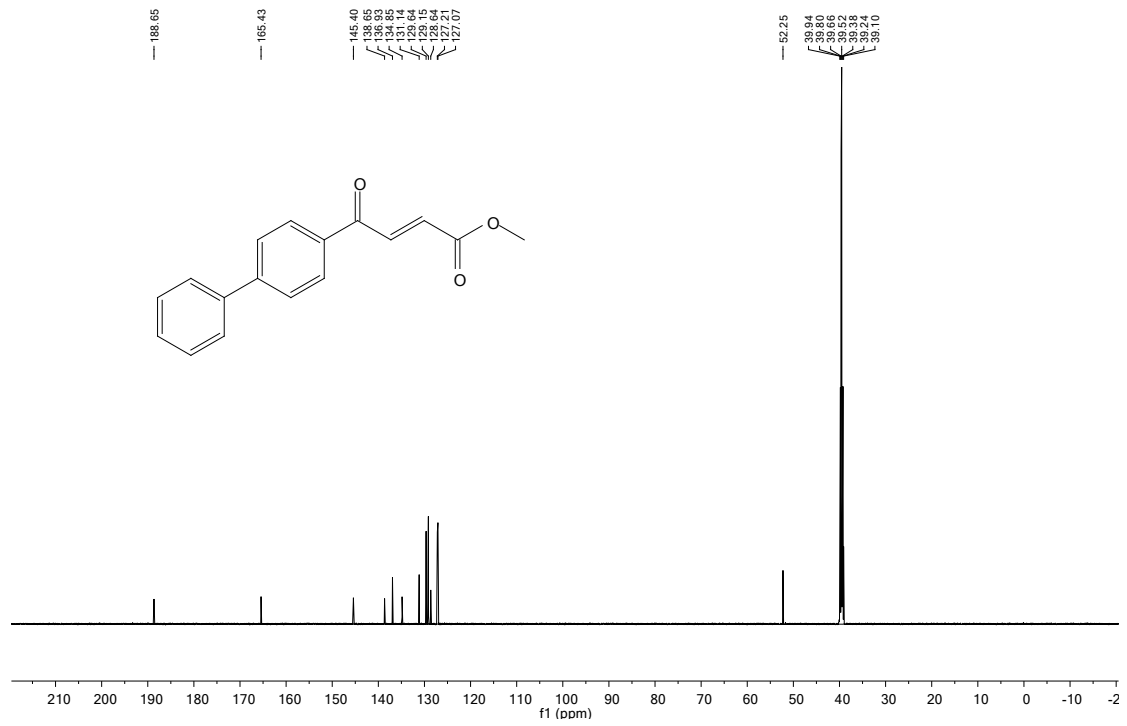
BRUKER AVANCE III 13C-NMR 8j IN DMSO-d6



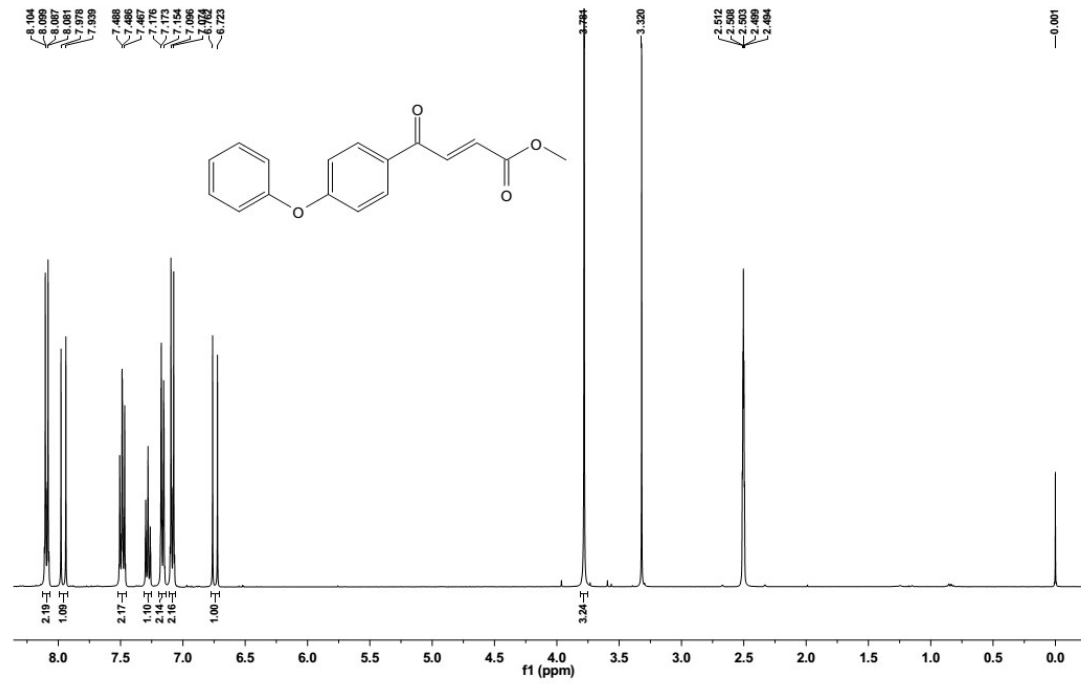
8k



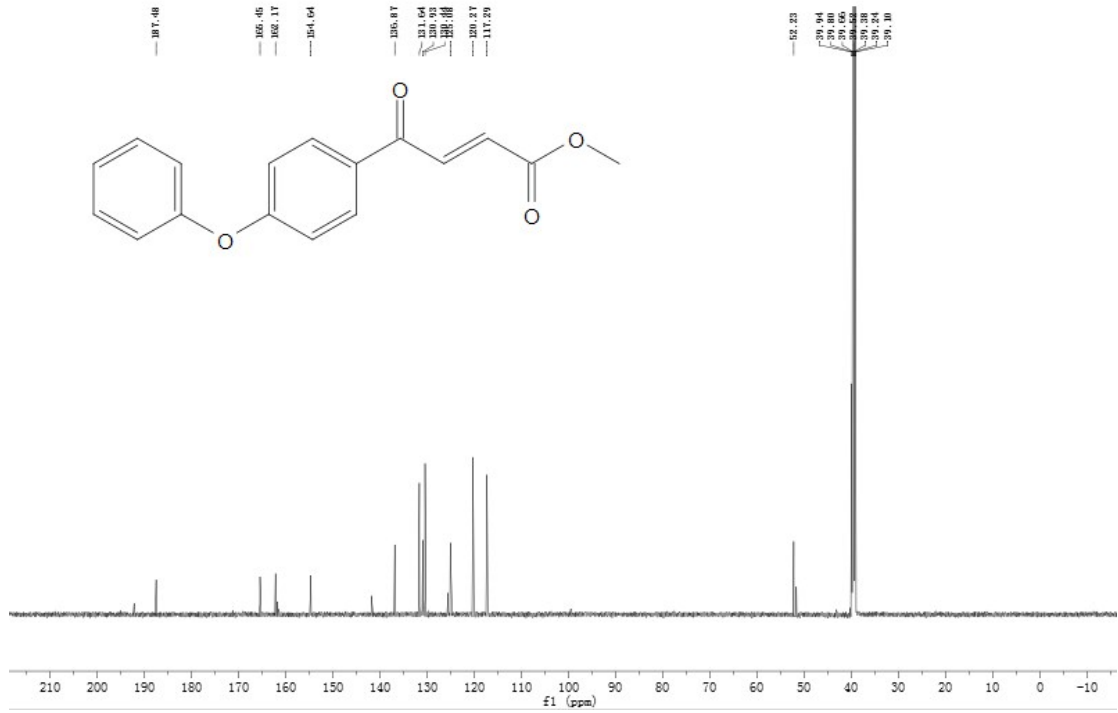
BRUKER AVANCE III 13C-NMR 8k IN DMSO-d6



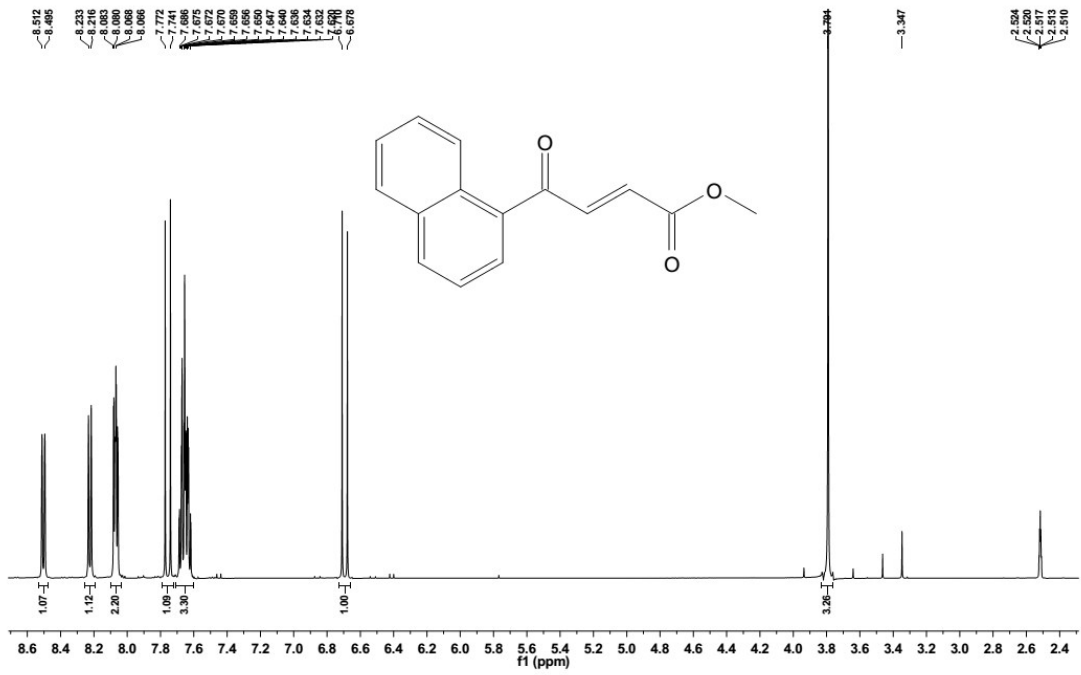
81



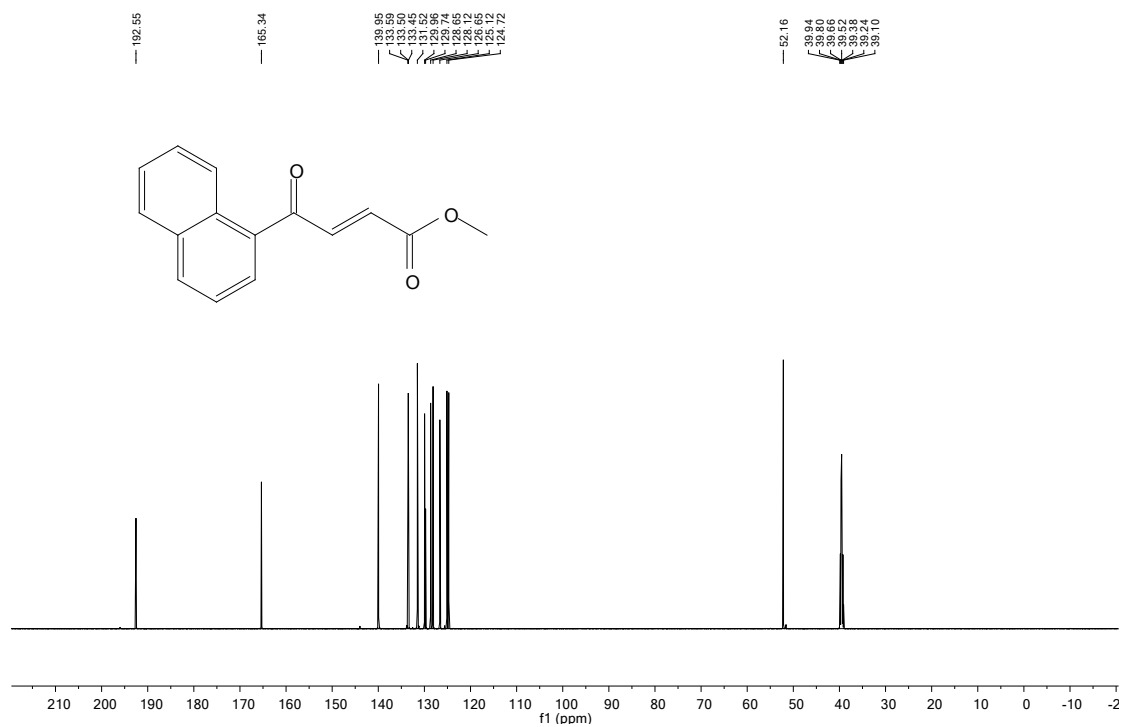
BRUKER AVANCE III 13C-NMR 81 IN DMSO-d6



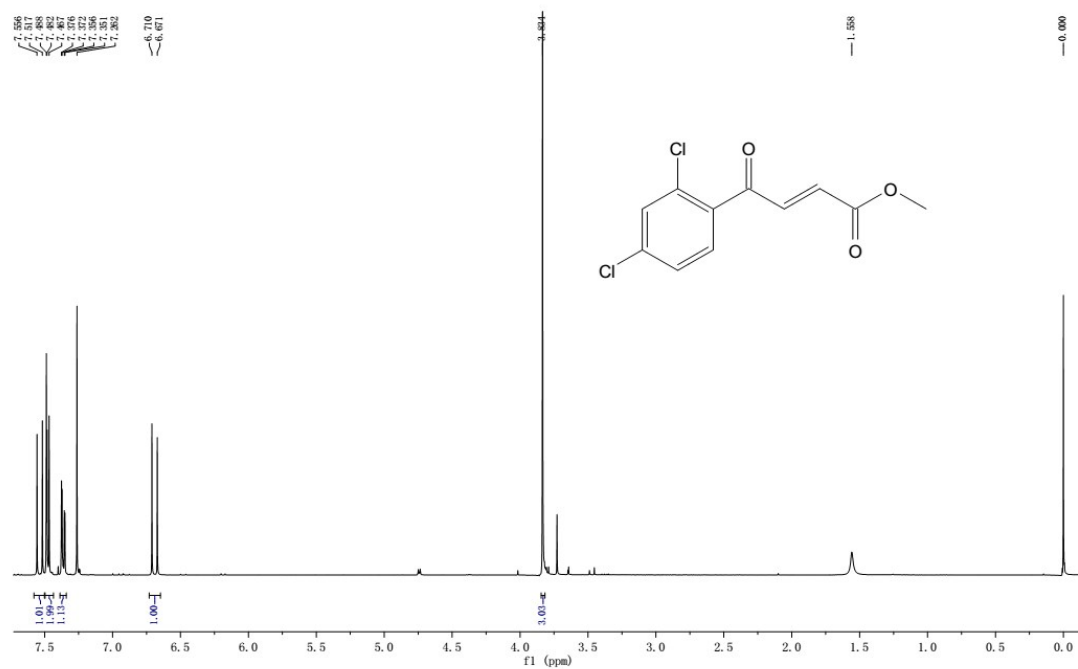
8m



BRUKER AVANCE III 13C-NMR 8m IN DMSO-d6



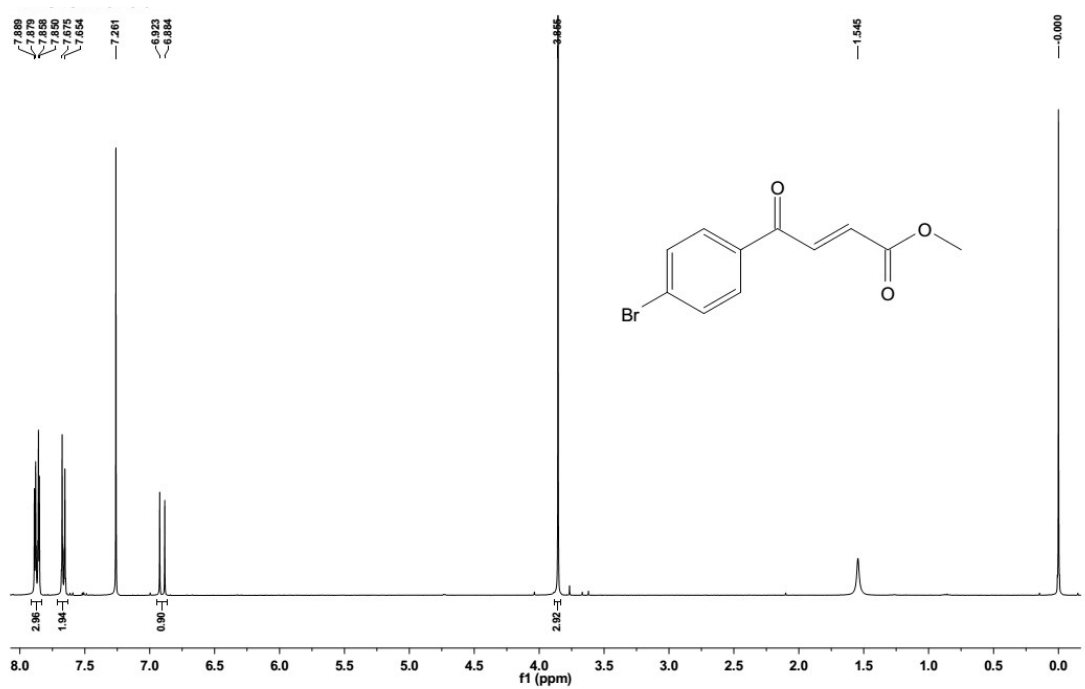
8n



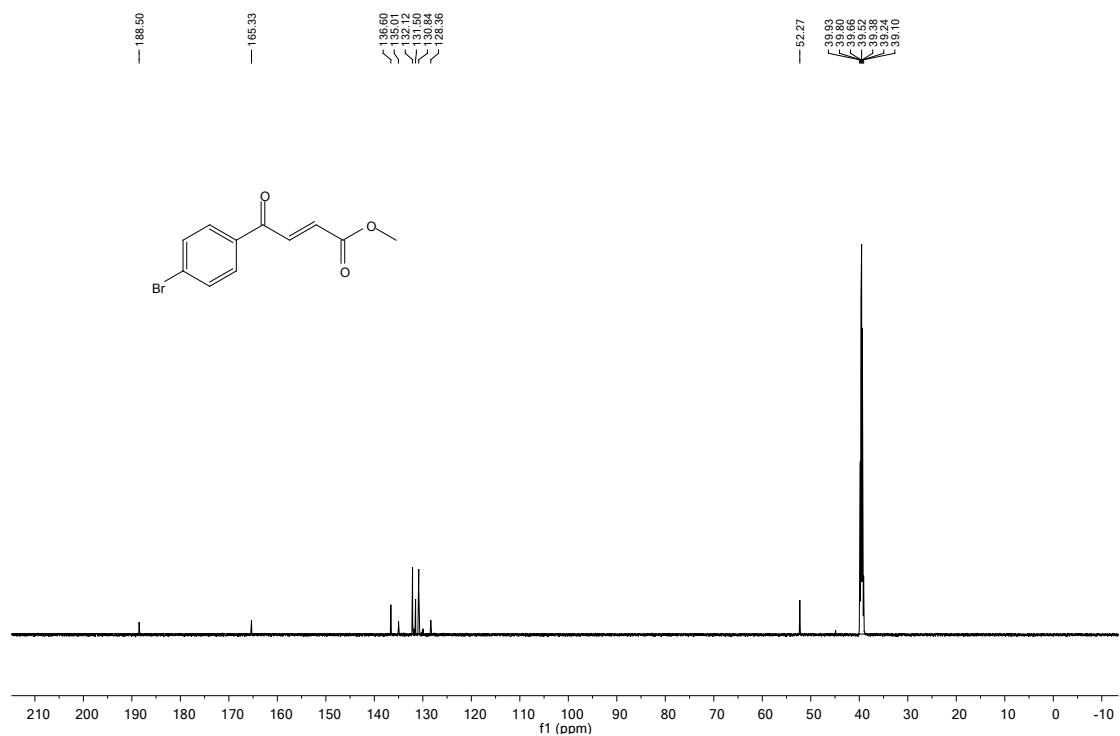
8o



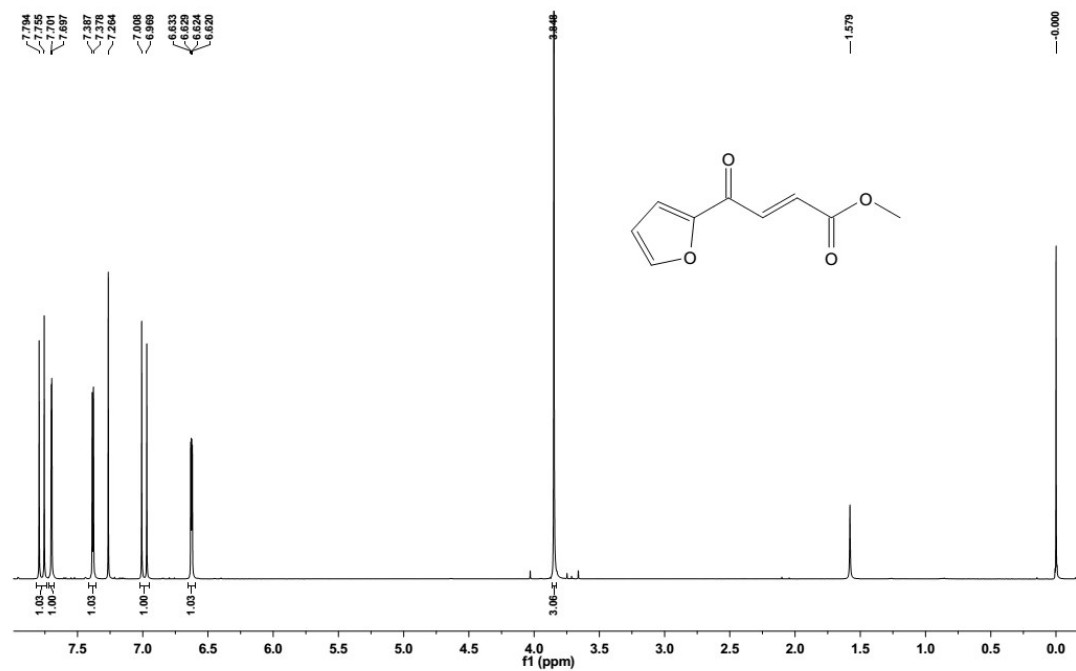
8p



BRUKER AVANCE III 13C-NMR 8p IN DMSO-d6



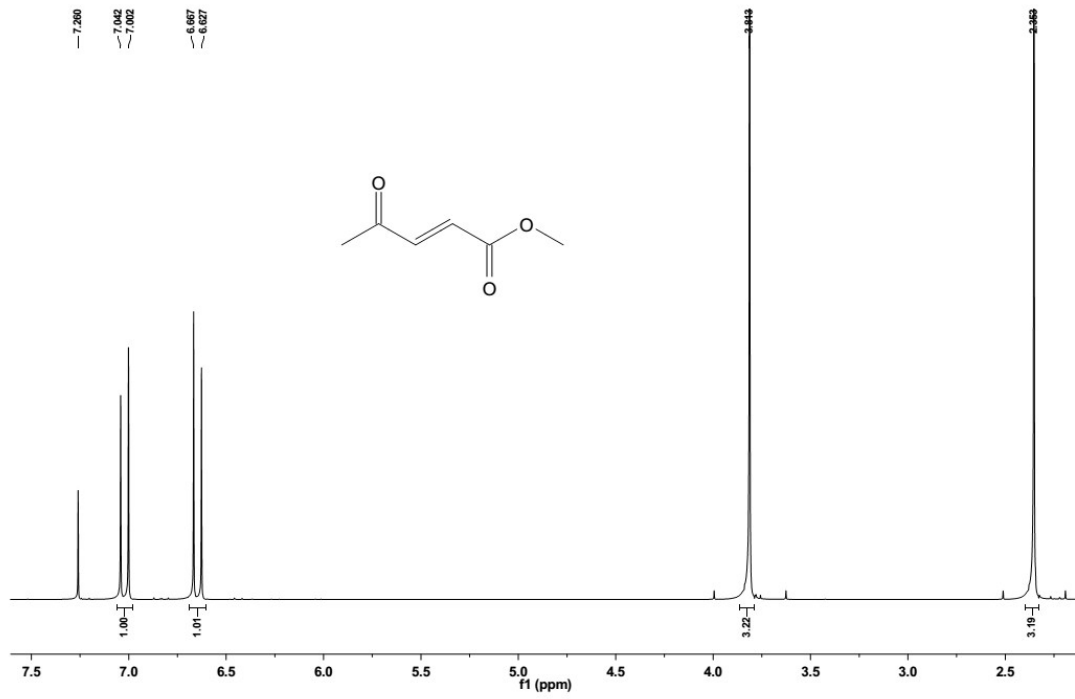
8q



BRUKER AVANCE III 13C-NMR 8q IN DMSO-d6



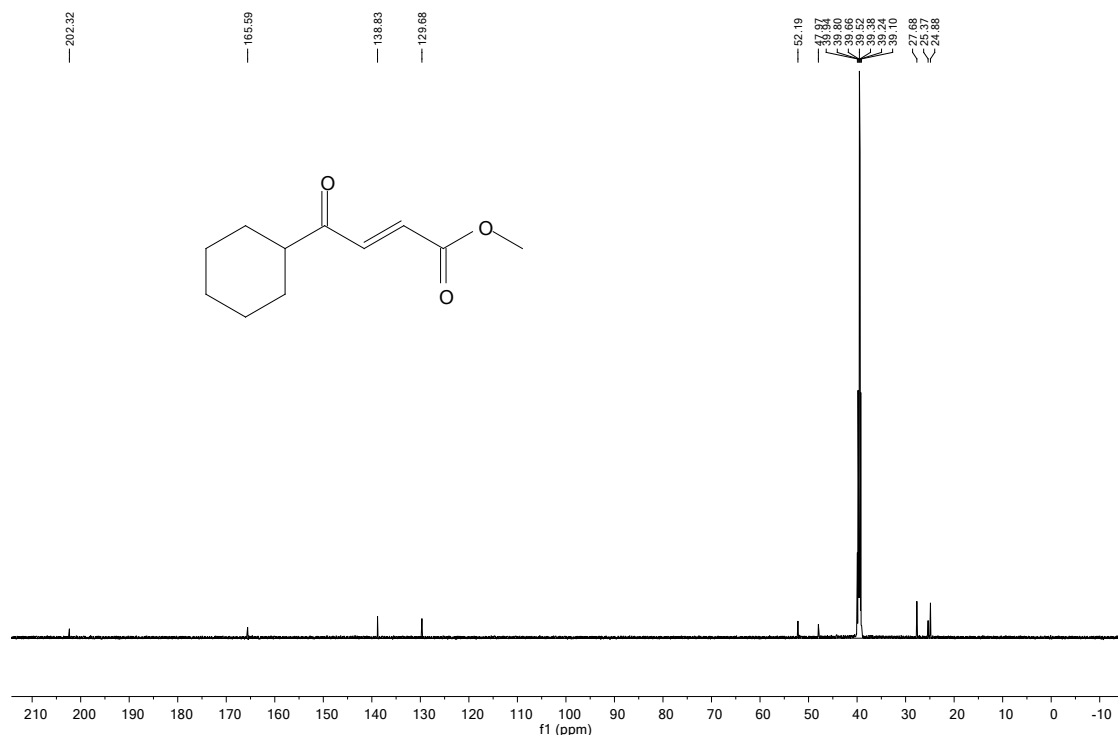
8r



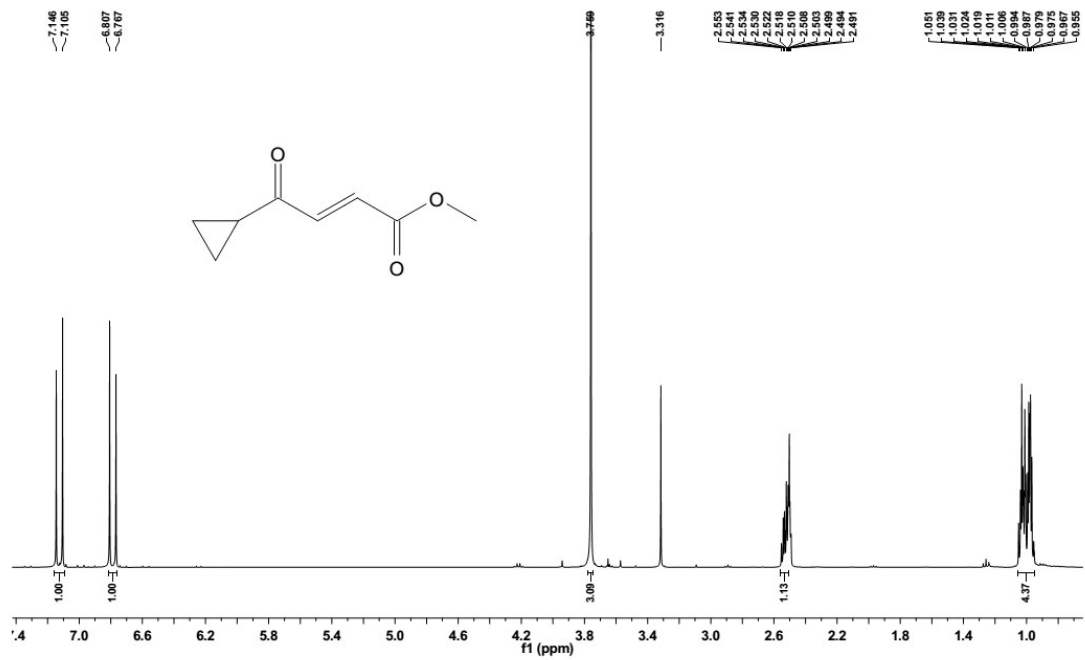
8s



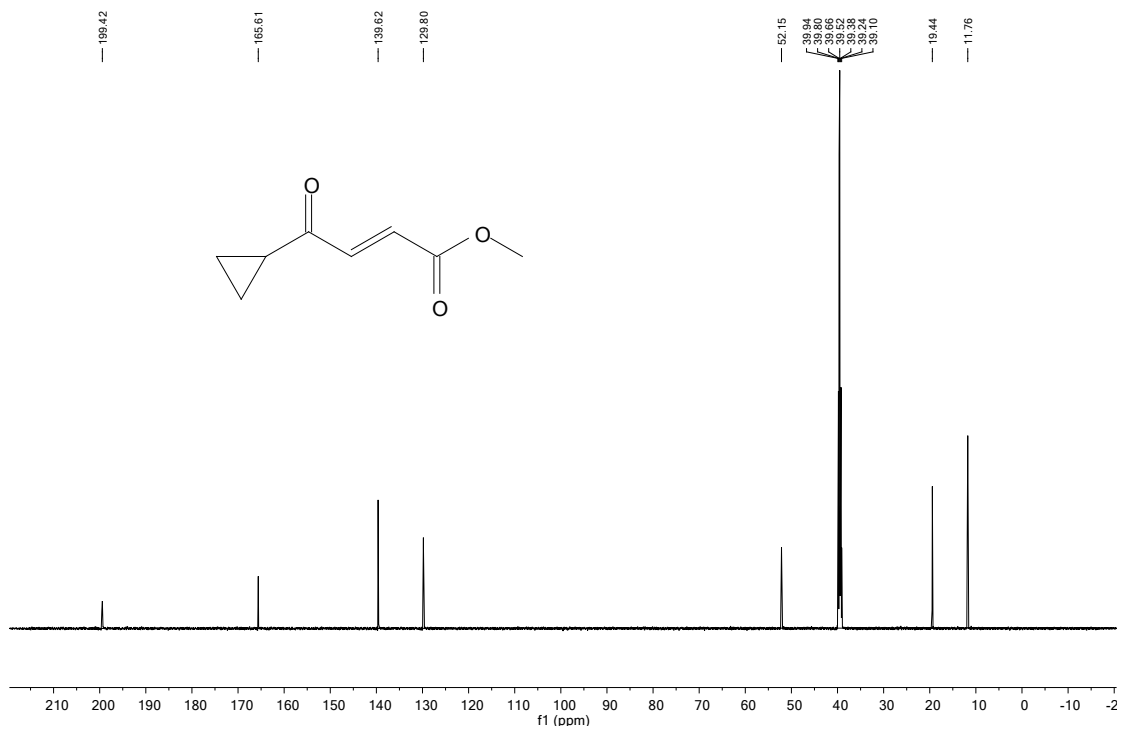
BRUKER AVANCE III 13C-NMR 8S IN DMSO-d₆



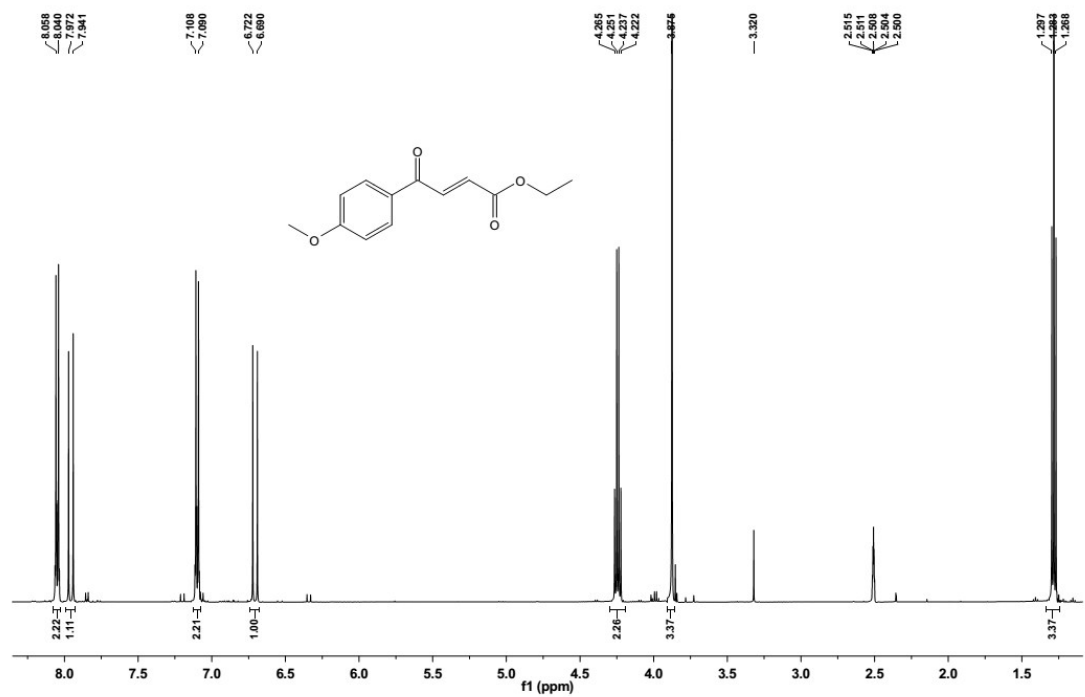
8t



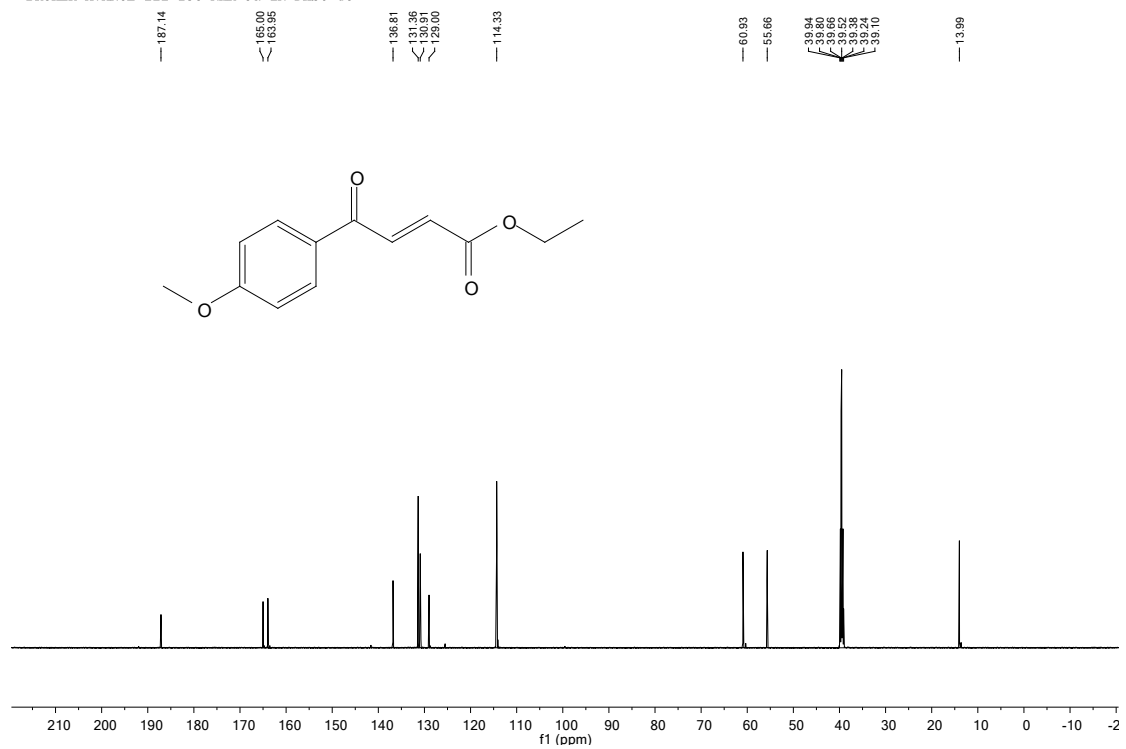
BRUKER AVANCE III 13C-NMR 8t IN DMSO-d6



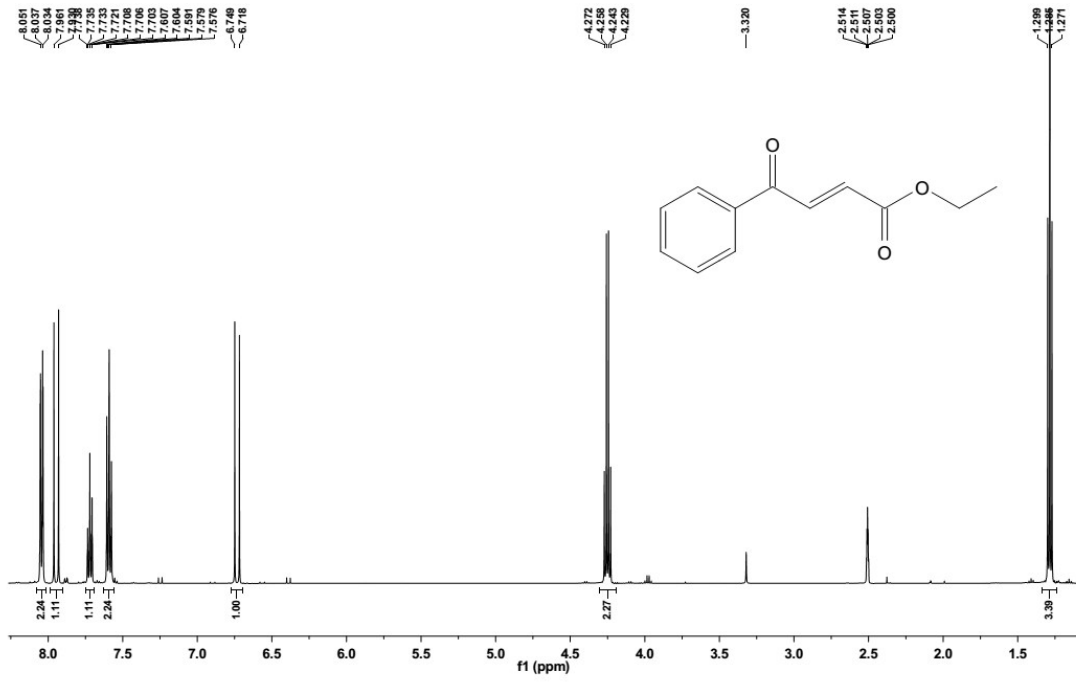
9a



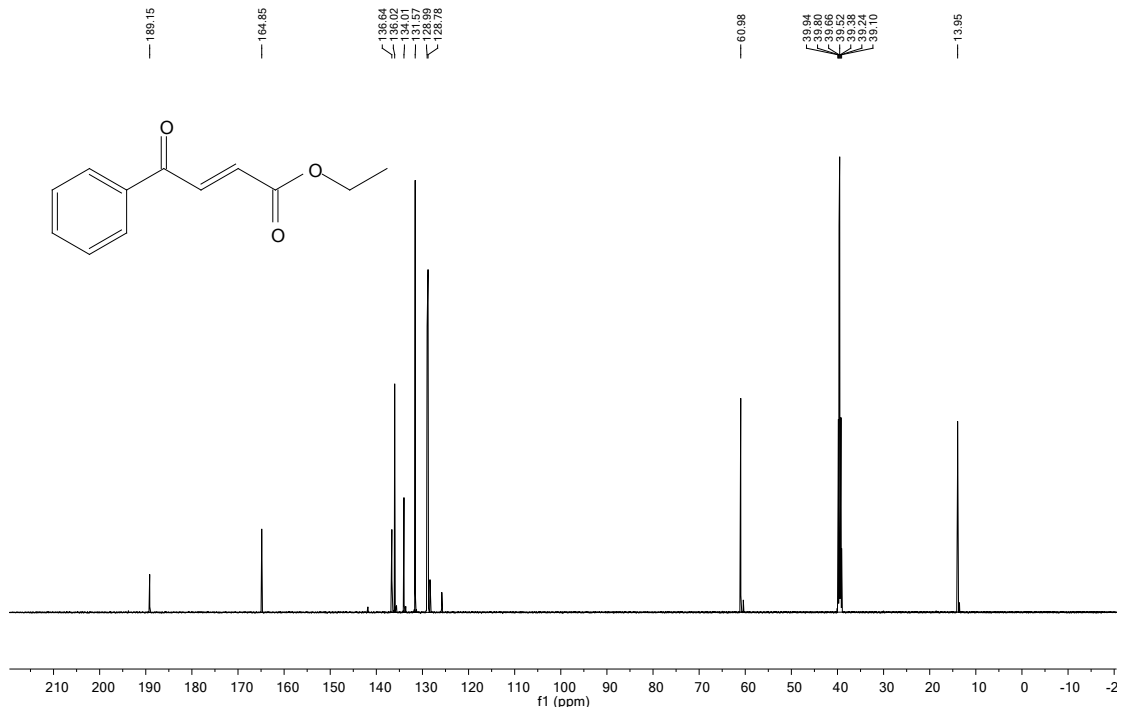
BRUKER AVANCE III 13C-NMR 9a IN DMSO-d6



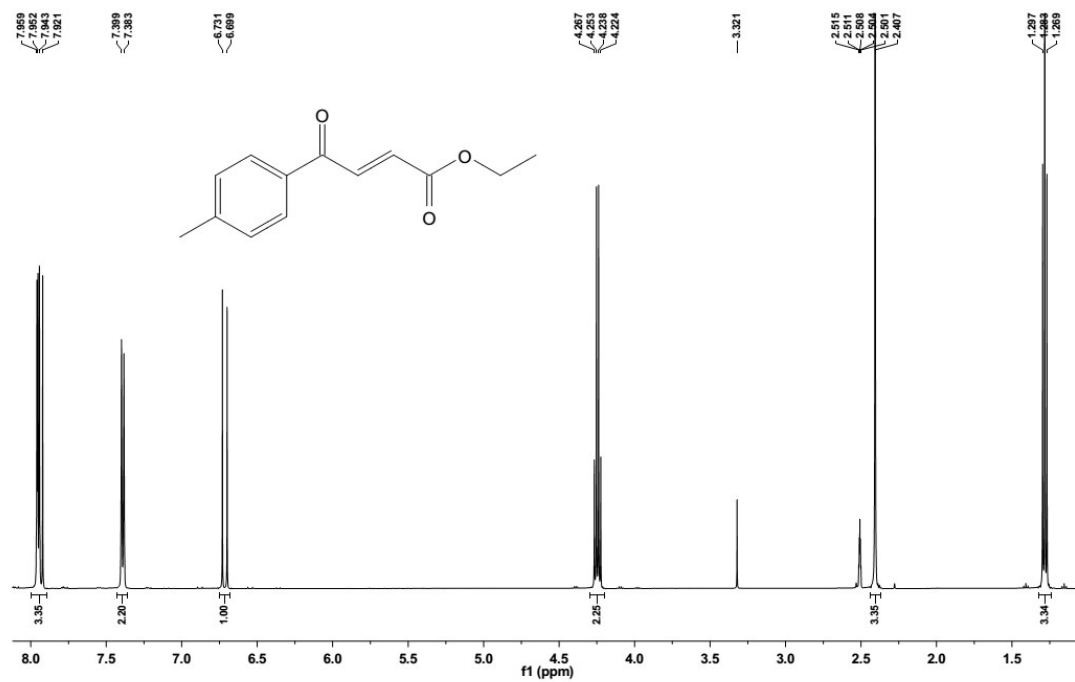
9b



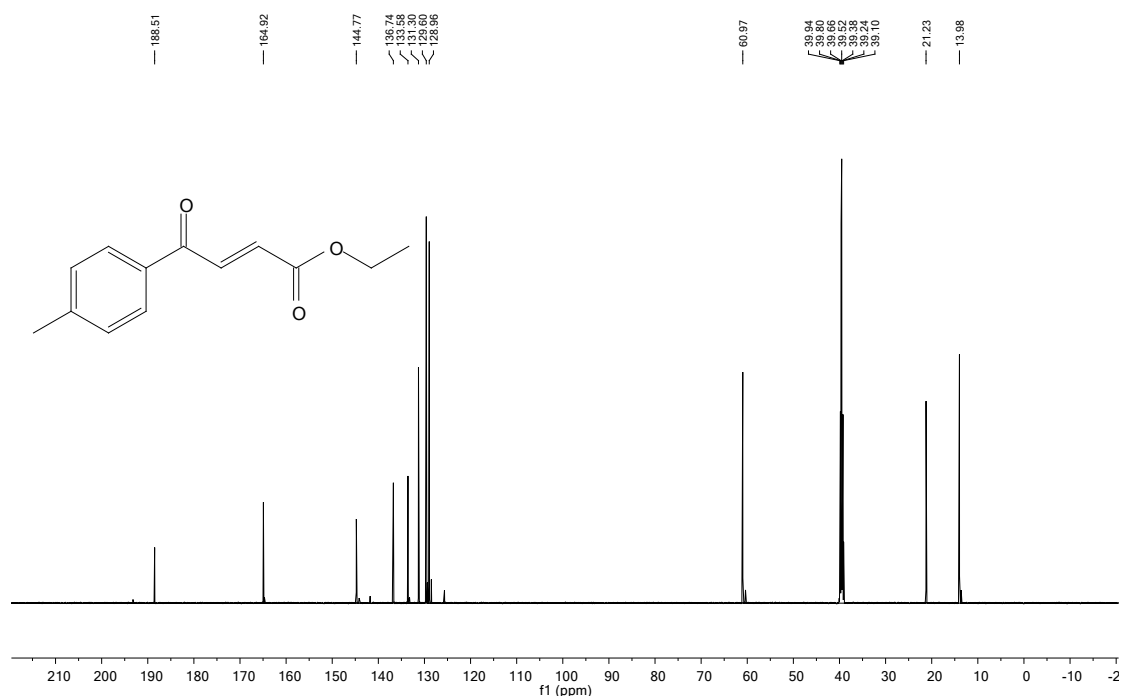
BRUKER AVANCE III 13C-NMR 9b IN DMSO-d6



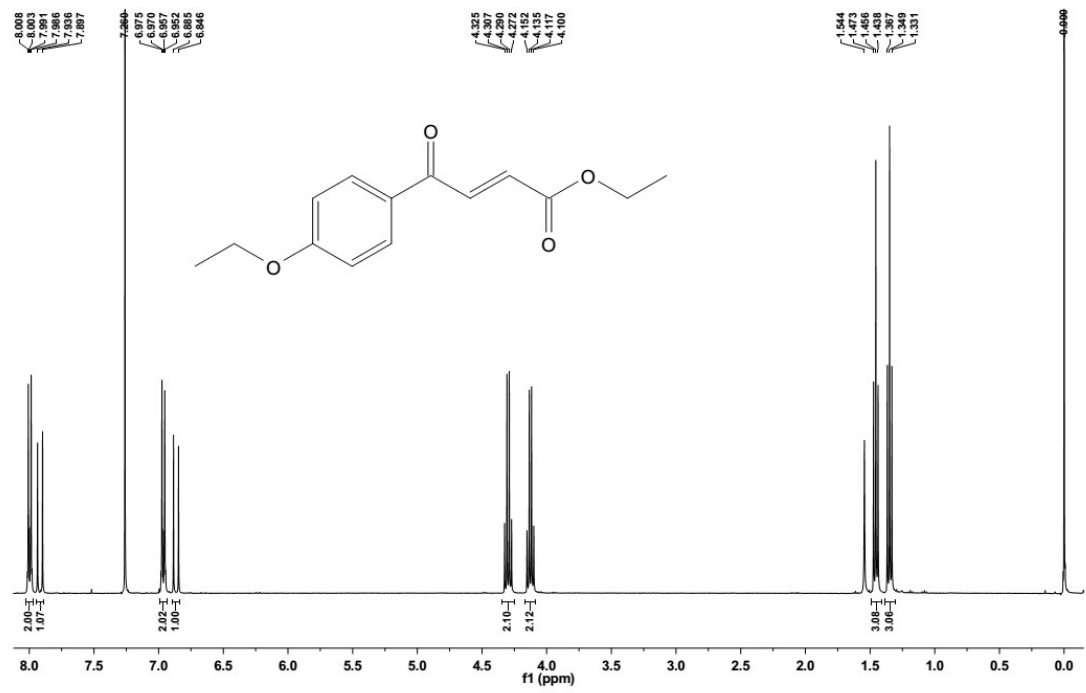
9c



BRUKER AVANCE III 13C-NMR 9c IN DMSO-d6



9d

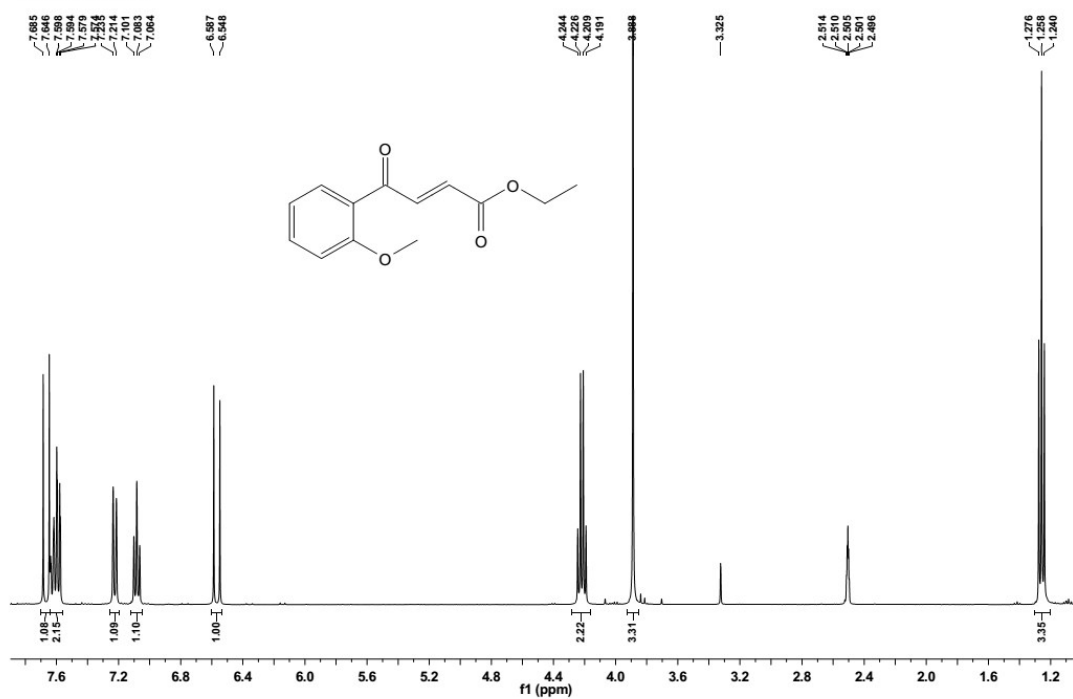


BRUKER AVANCE III 13C-NMR 9d IN DMSO-d6

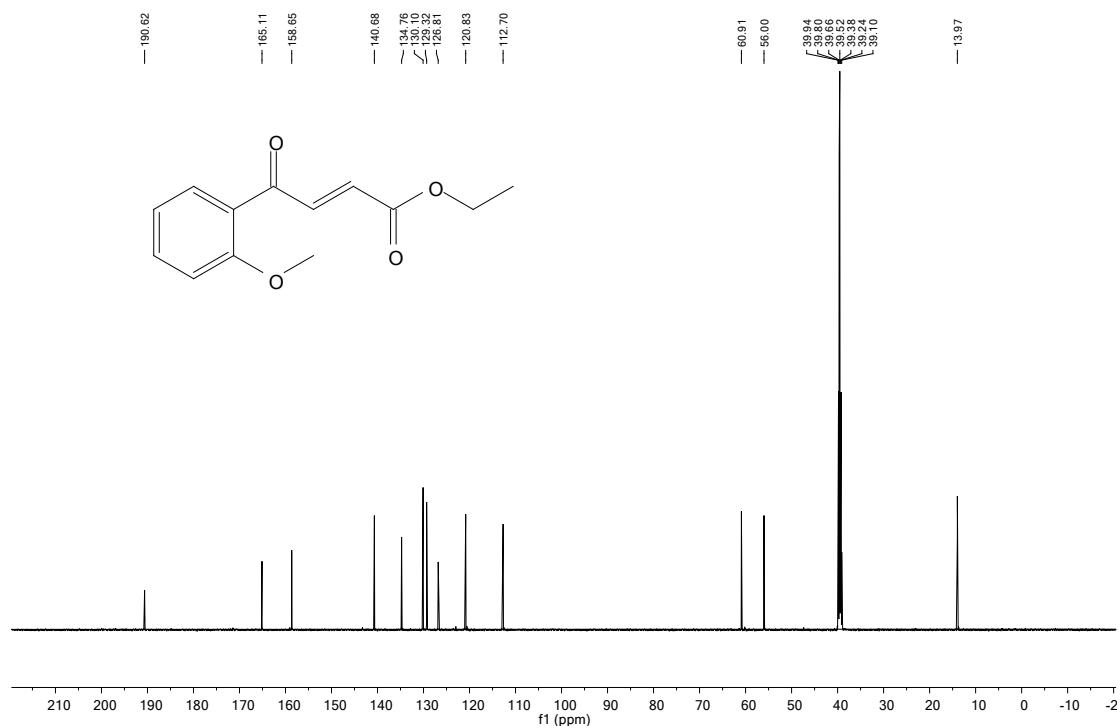


9e

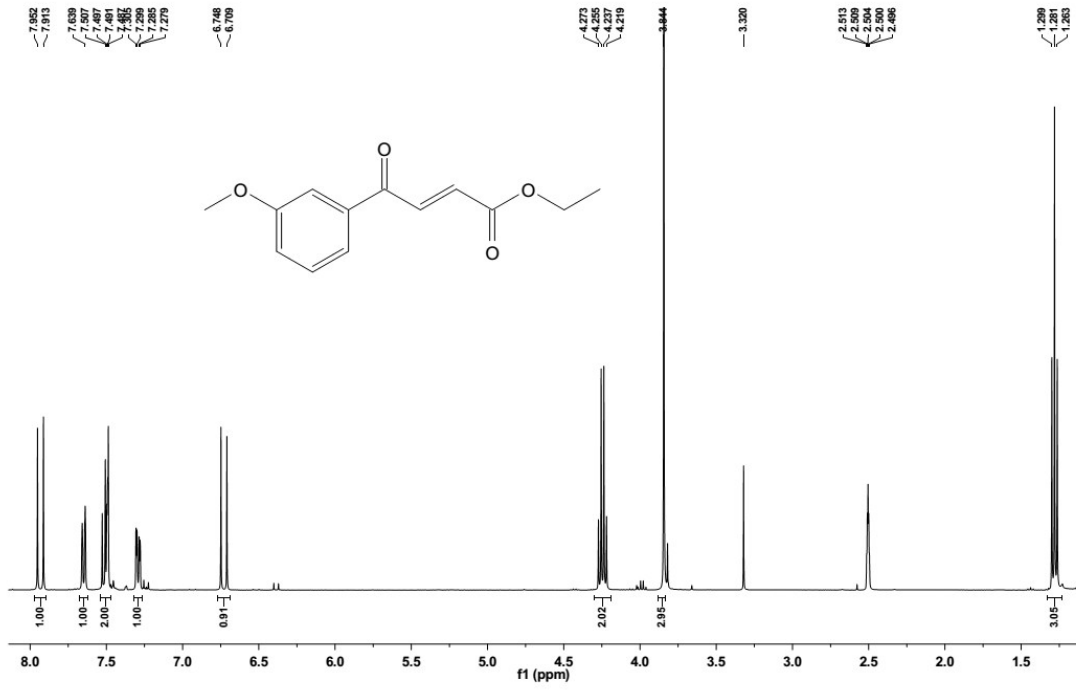
Bruker AVANCE III 400 20131014_XCL13E04
PROTON2 DMSO



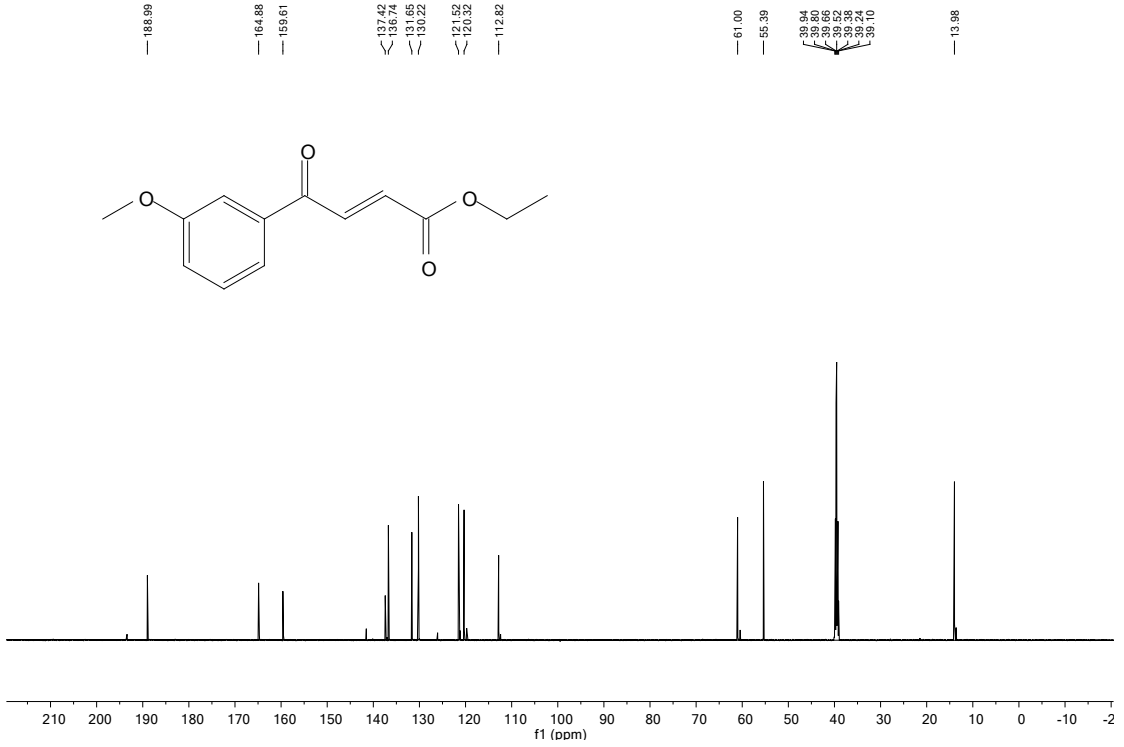
BRUKER AVANCE III 13C-NMR 9e IN DMSO-d6



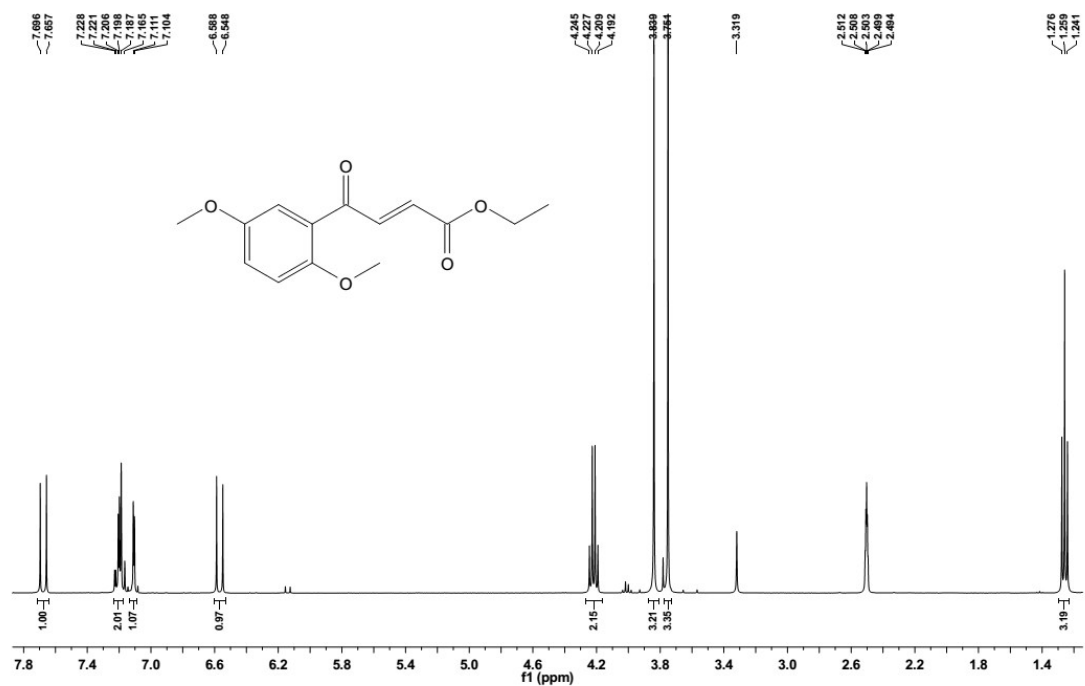
9f



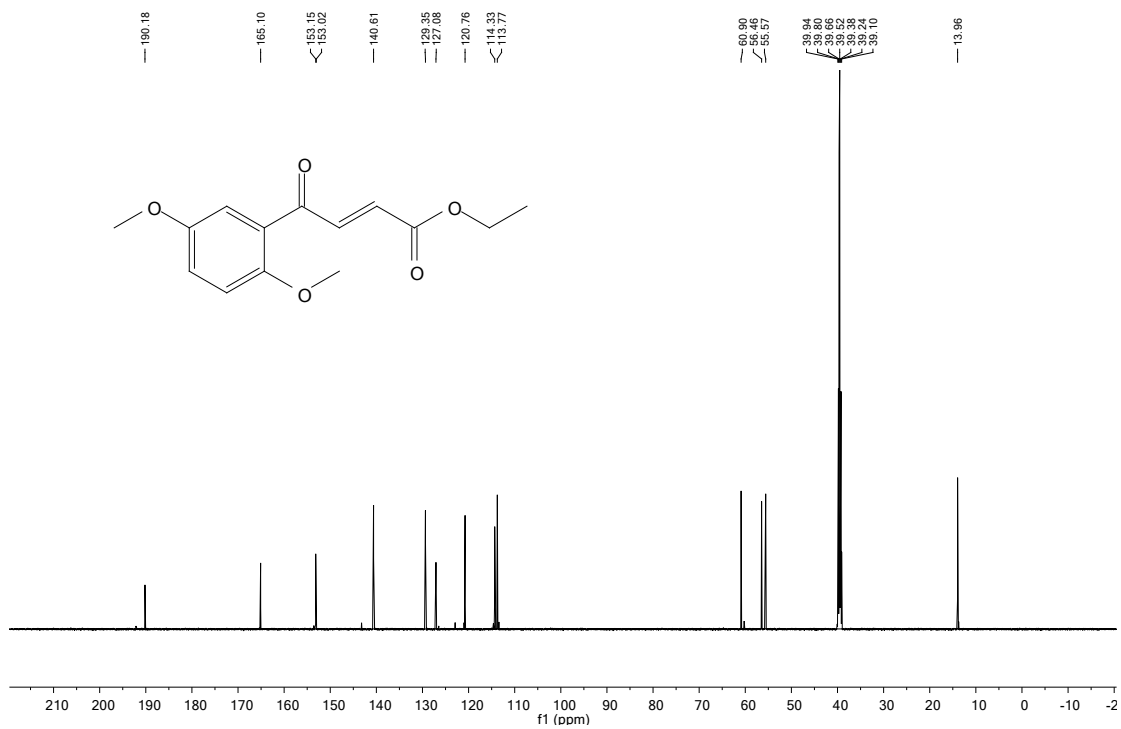
BRUKER AVANCE III 13C-NMR 9f IN DMSO-d6



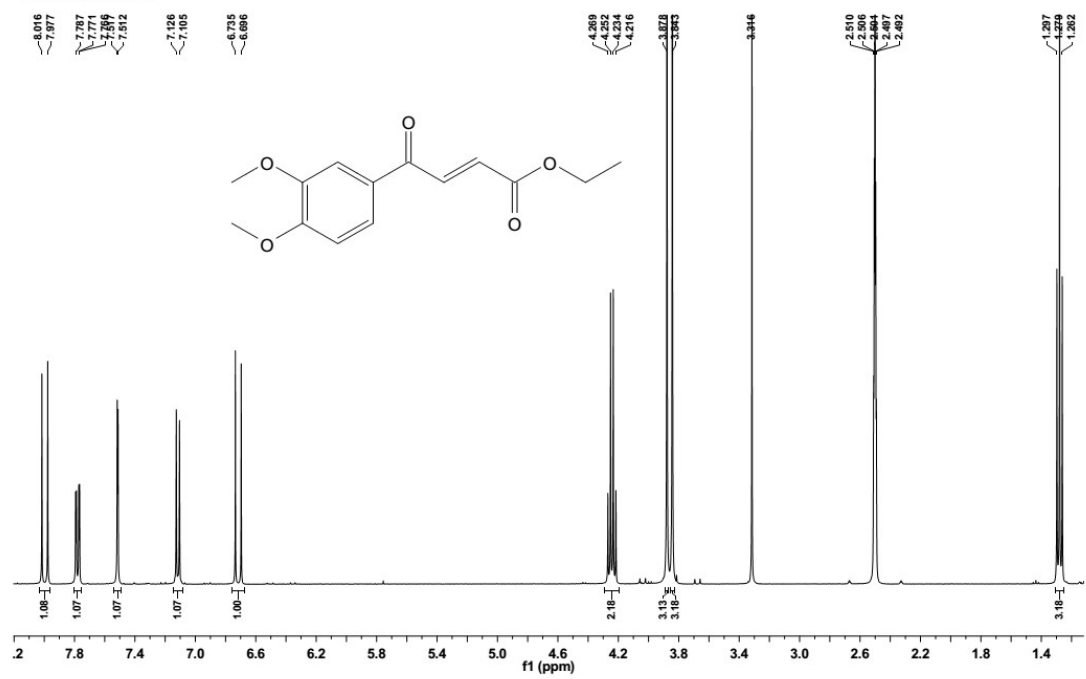
9g



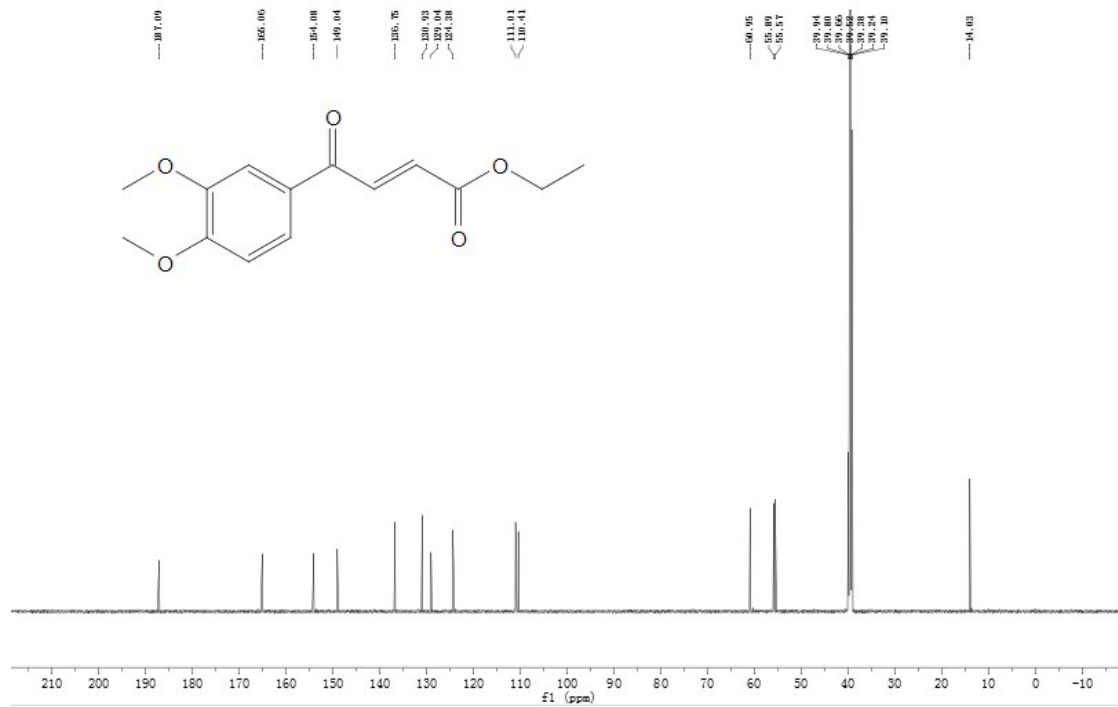
BRUKER AVANCE III 13C-NMR 9g IN DMSO-d6



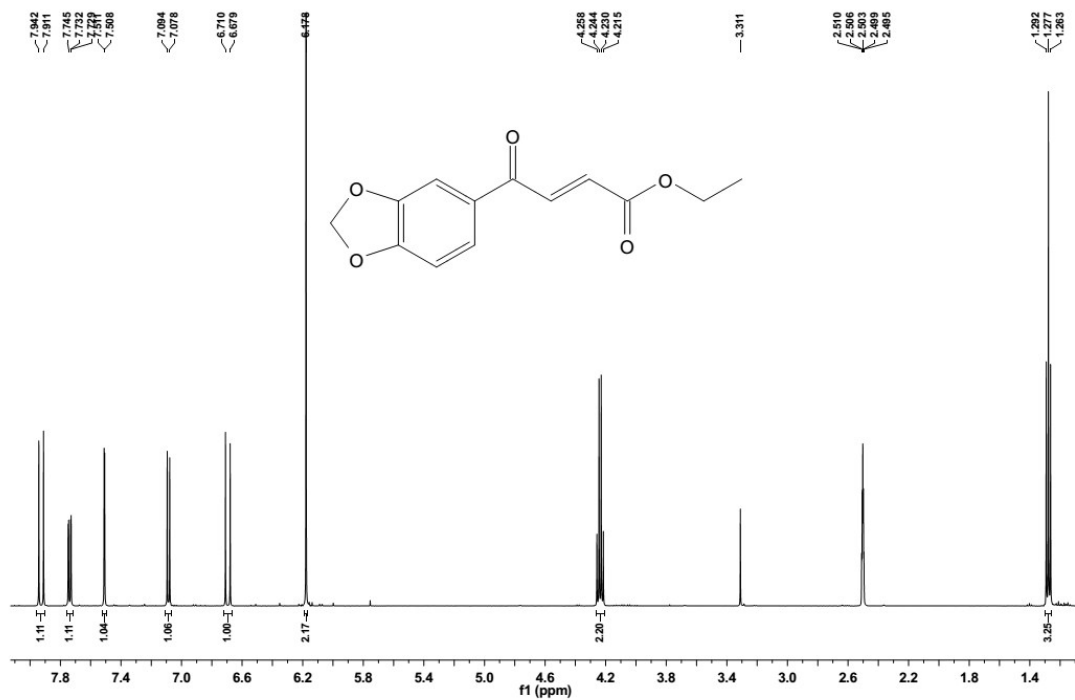
9h



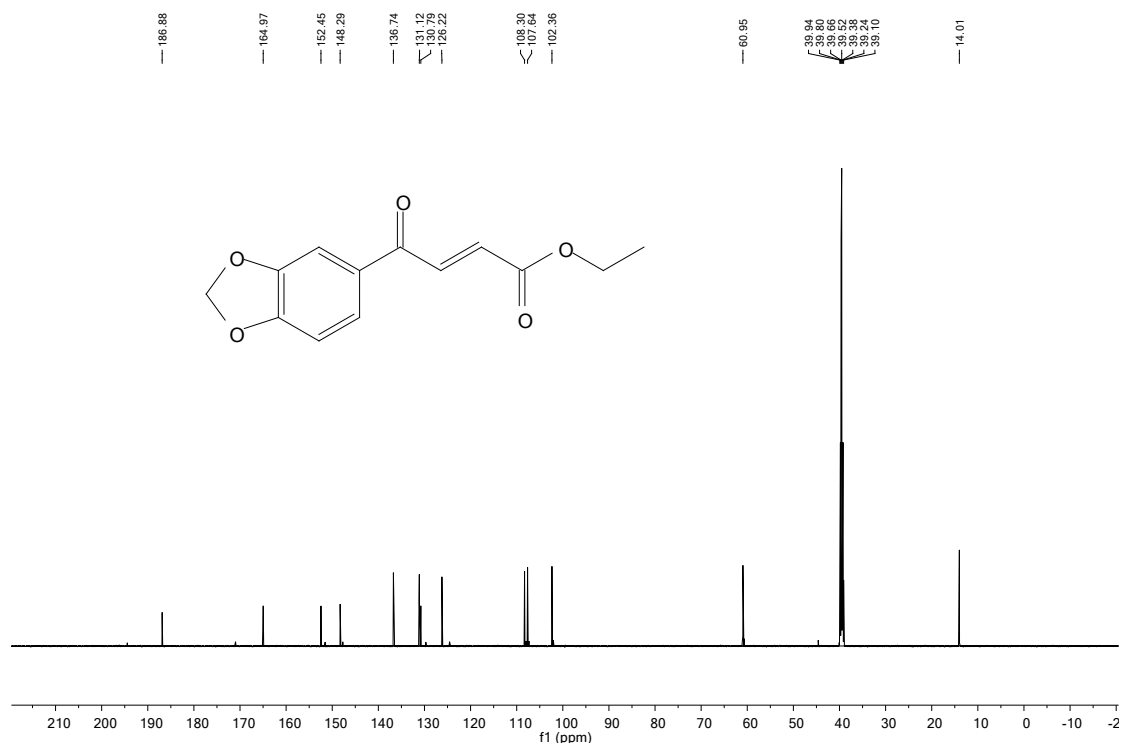
BRUKER AVANCE III 13C-NMR 9h IN DMSO-d6



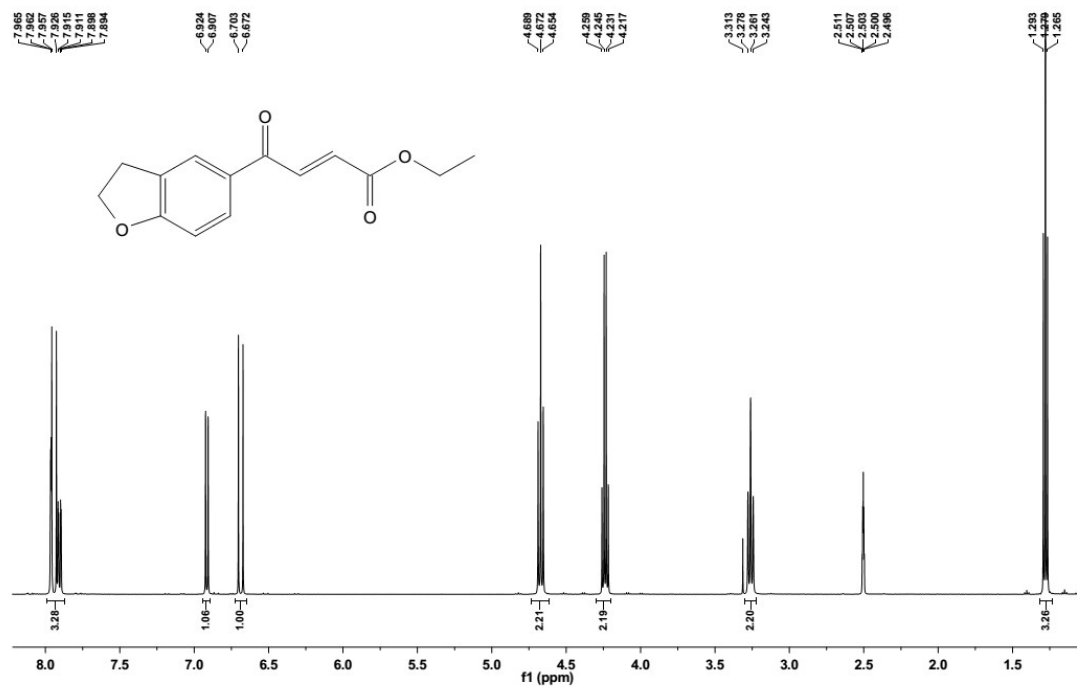
9i



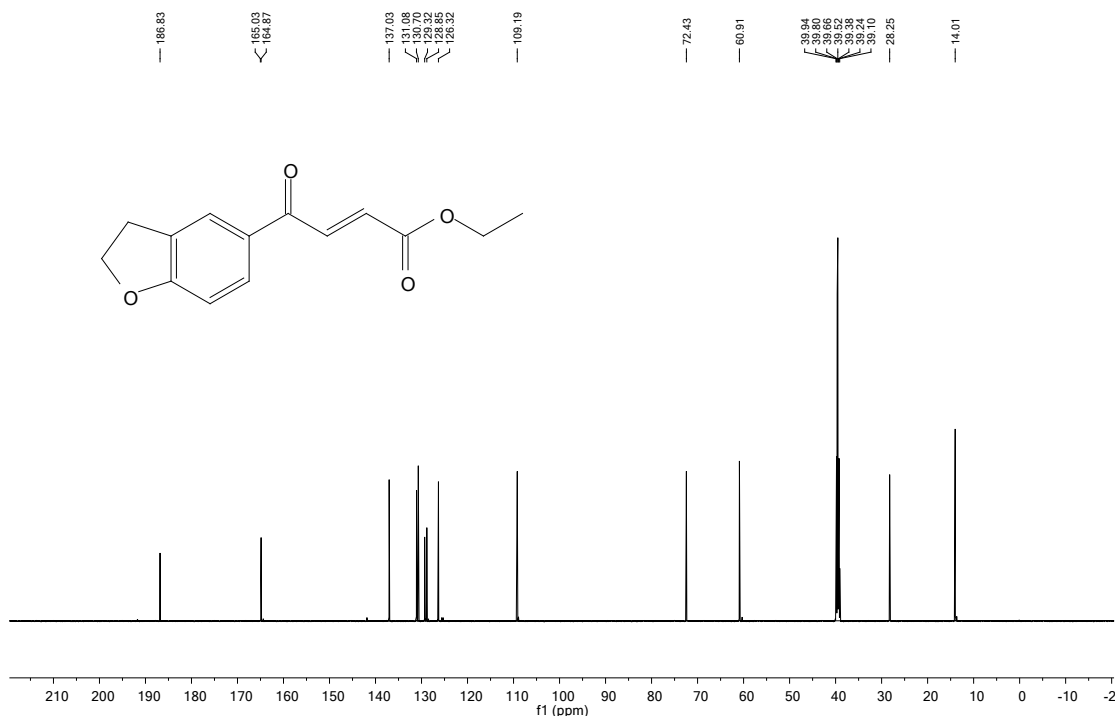
BRUKER AVANCE III 13C-NMR 9i IN DMSO-d6



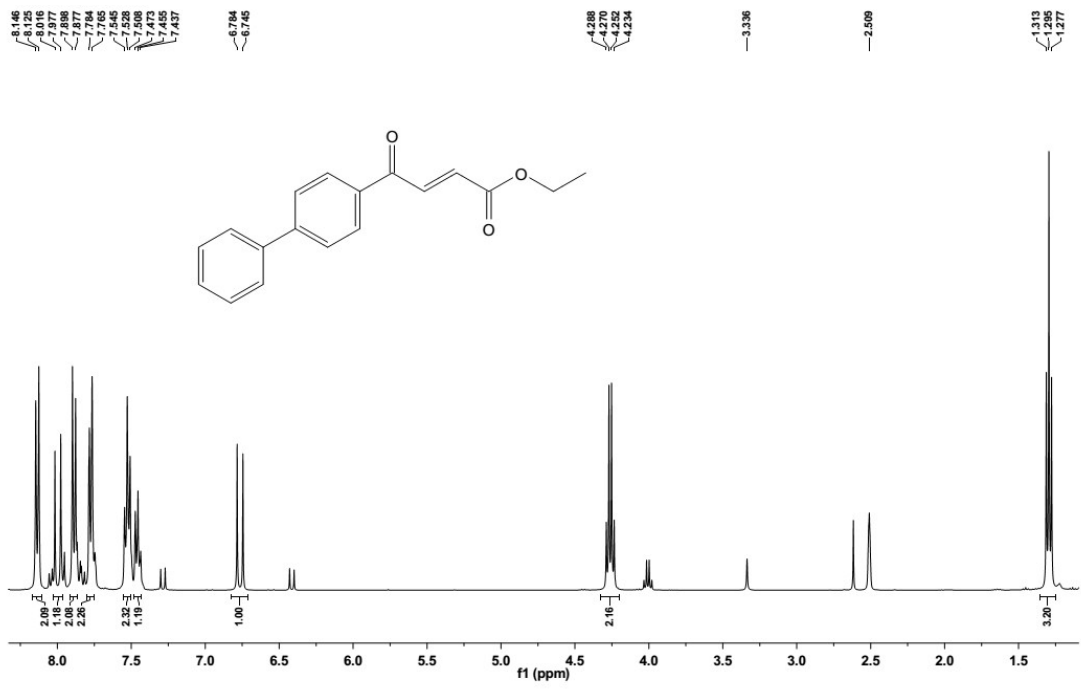
9j



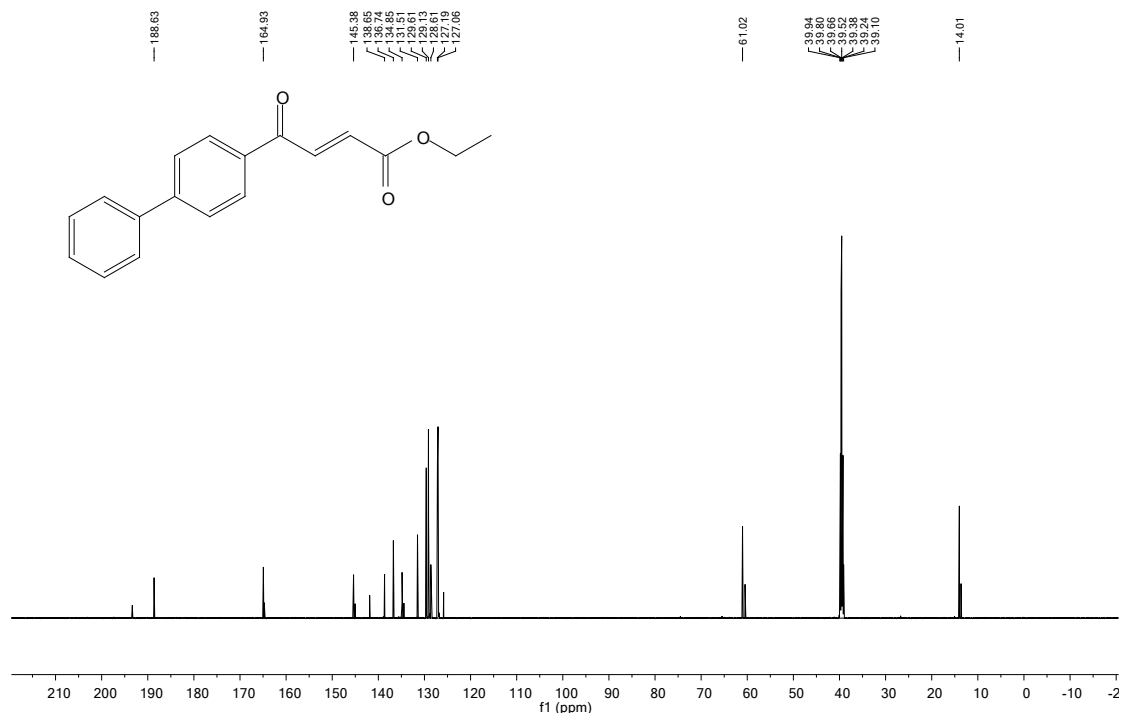
BRUKER AVANCE III 13C-NMR 9j IN DMSO-d6

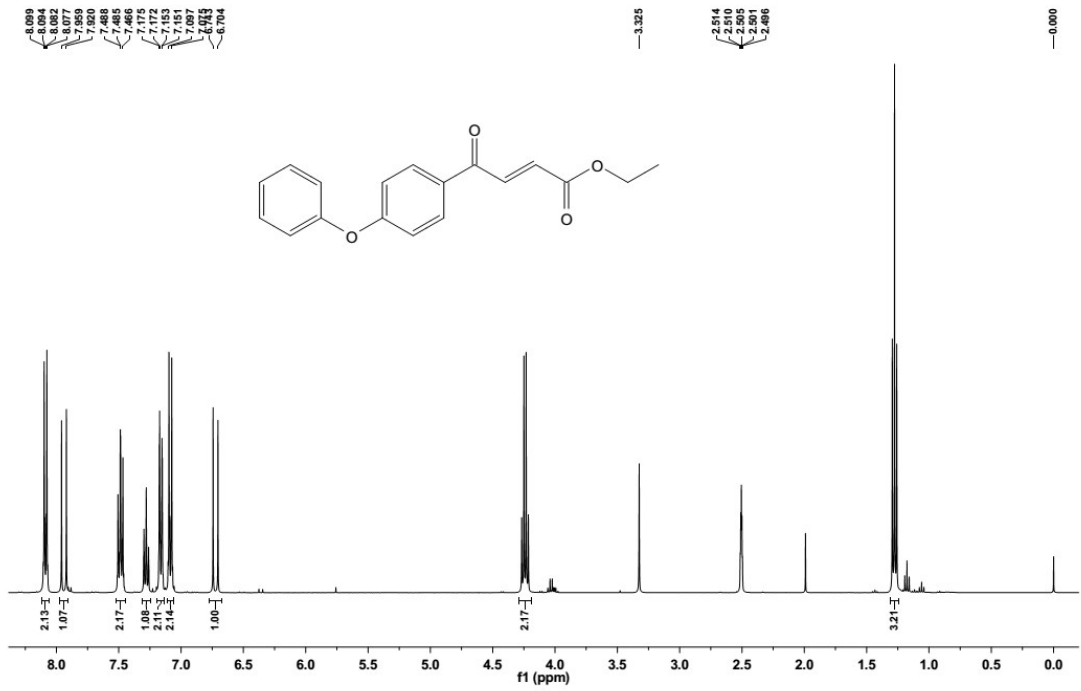


9k

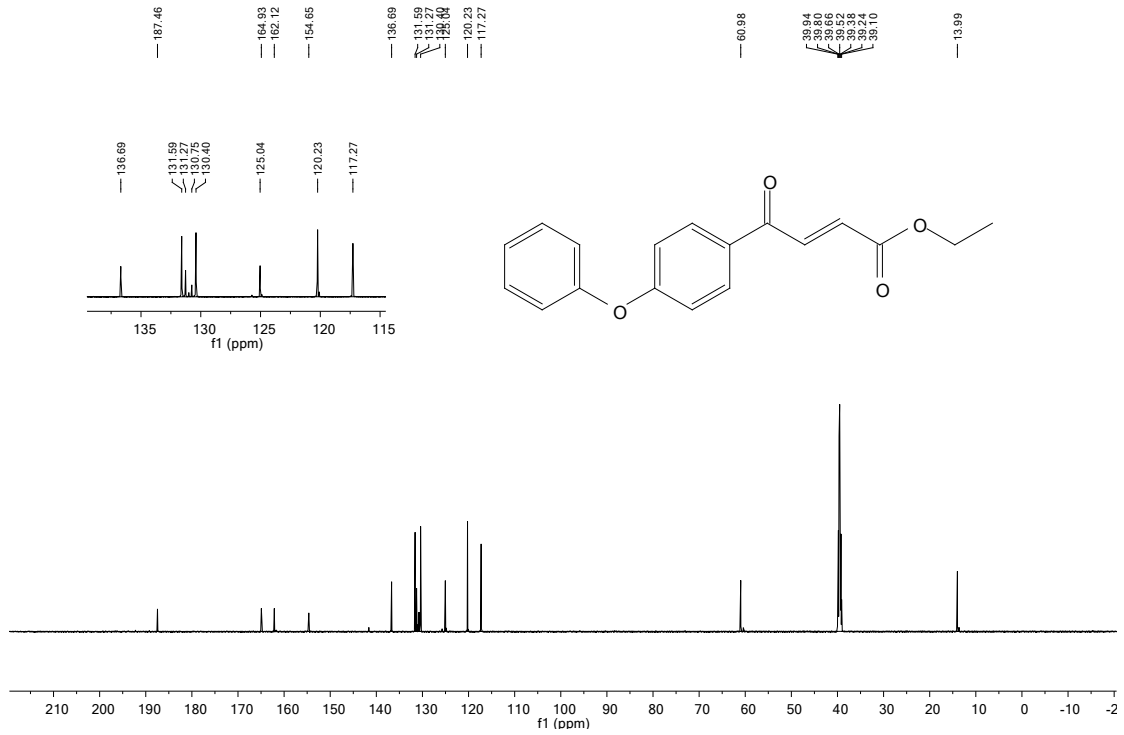


BRUKER AVANCE III 13C-NMR 9k IN DMSO-d6

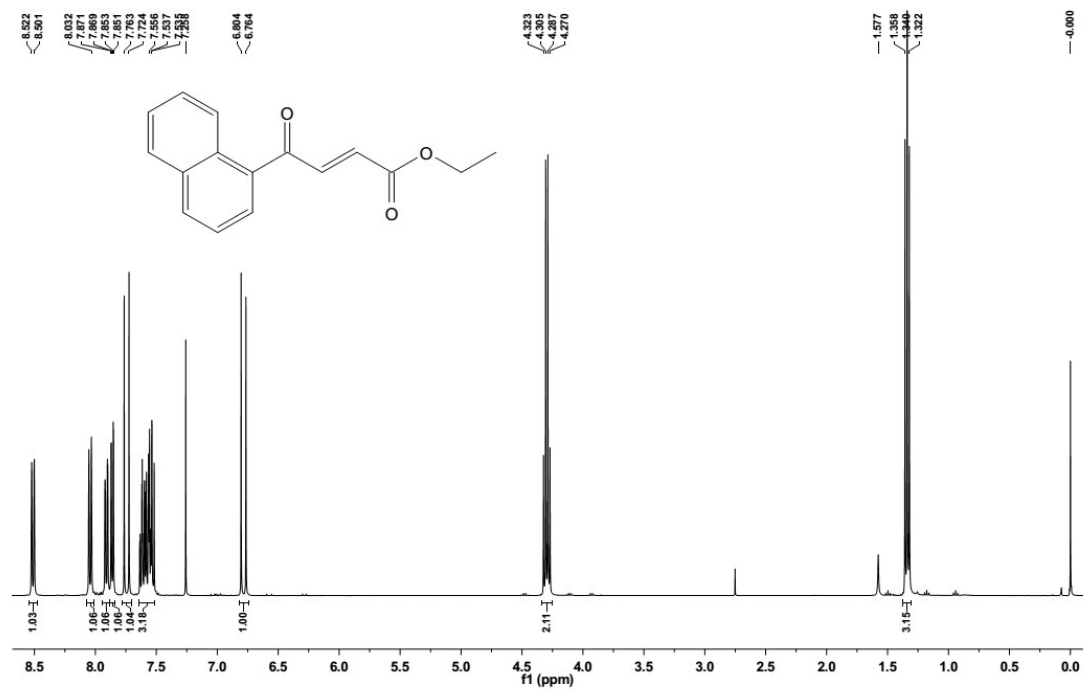




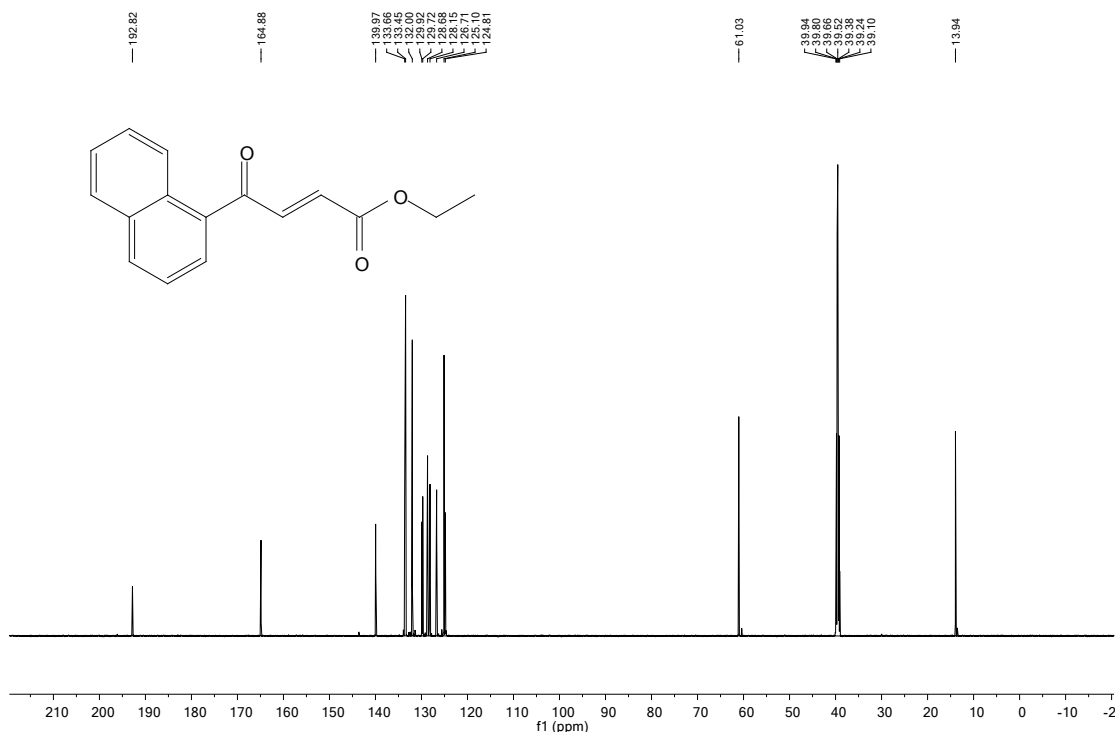
BRUKER AVANCE III 13C-NMR 91 IN DMSO-d₆



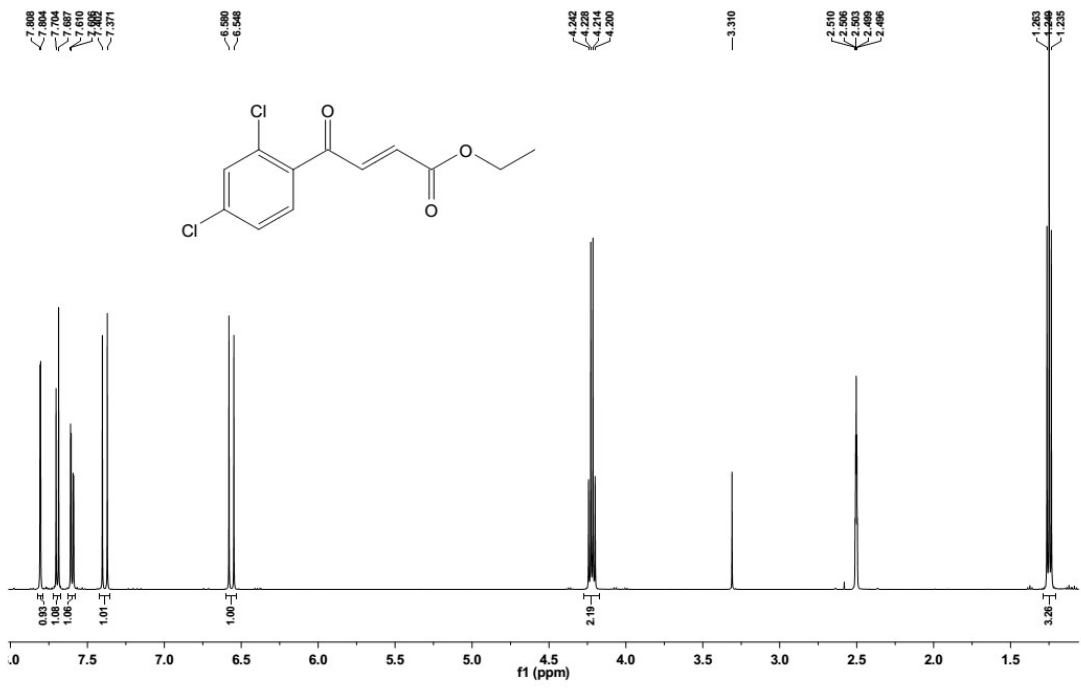
9m



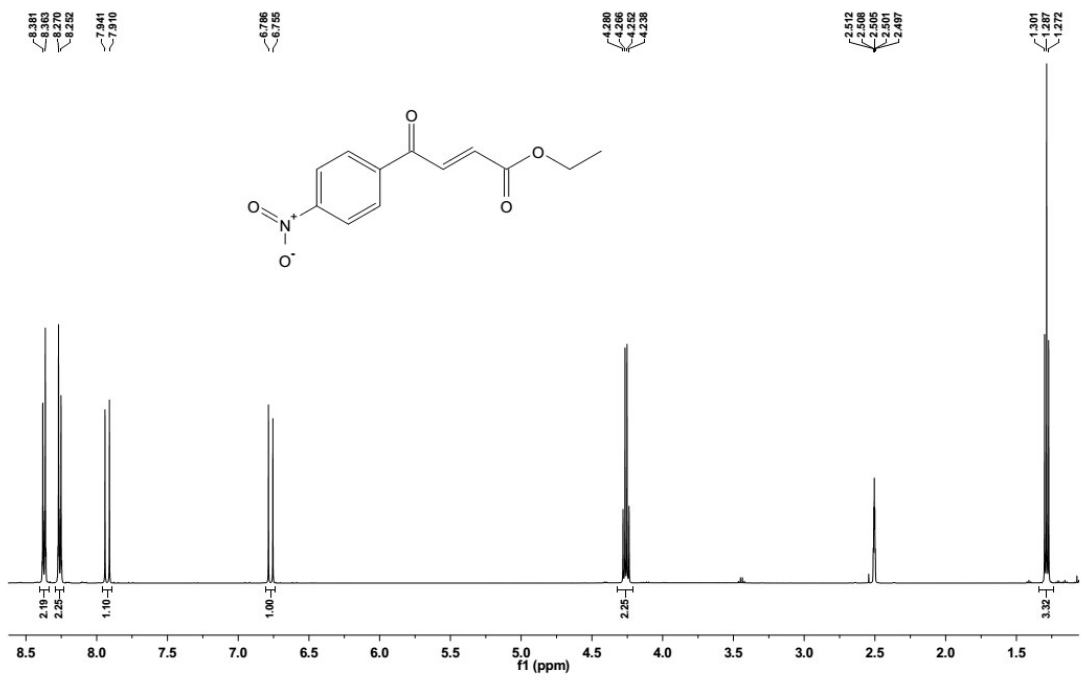
BRUKER AVANCE III 13C-NMR 9m IN DMSO-d6



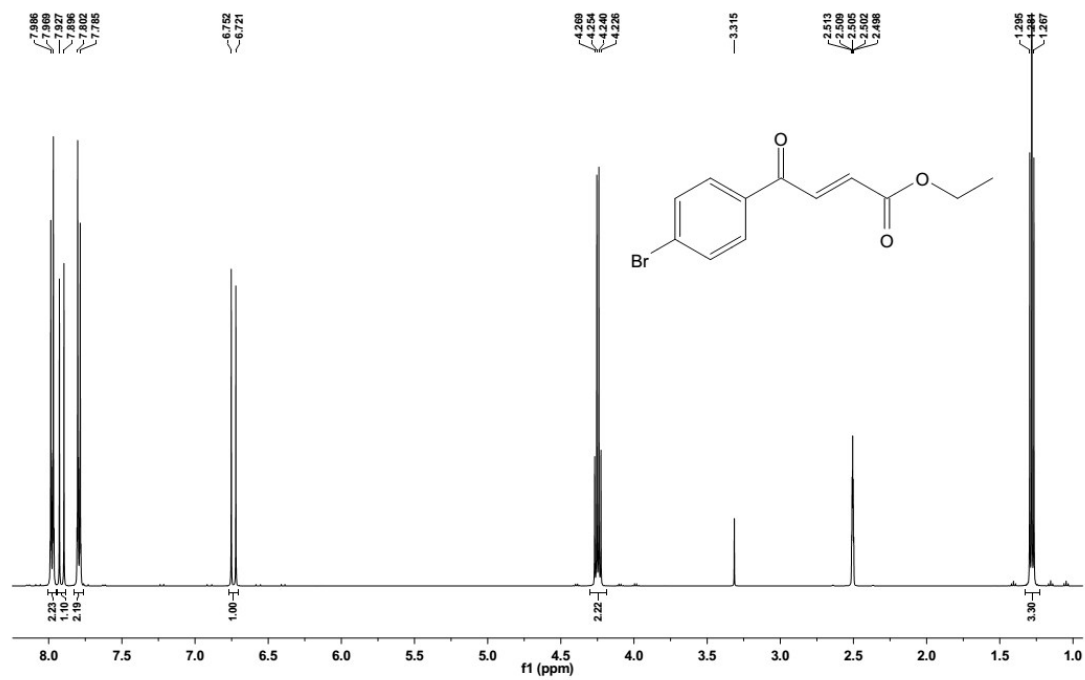
9n



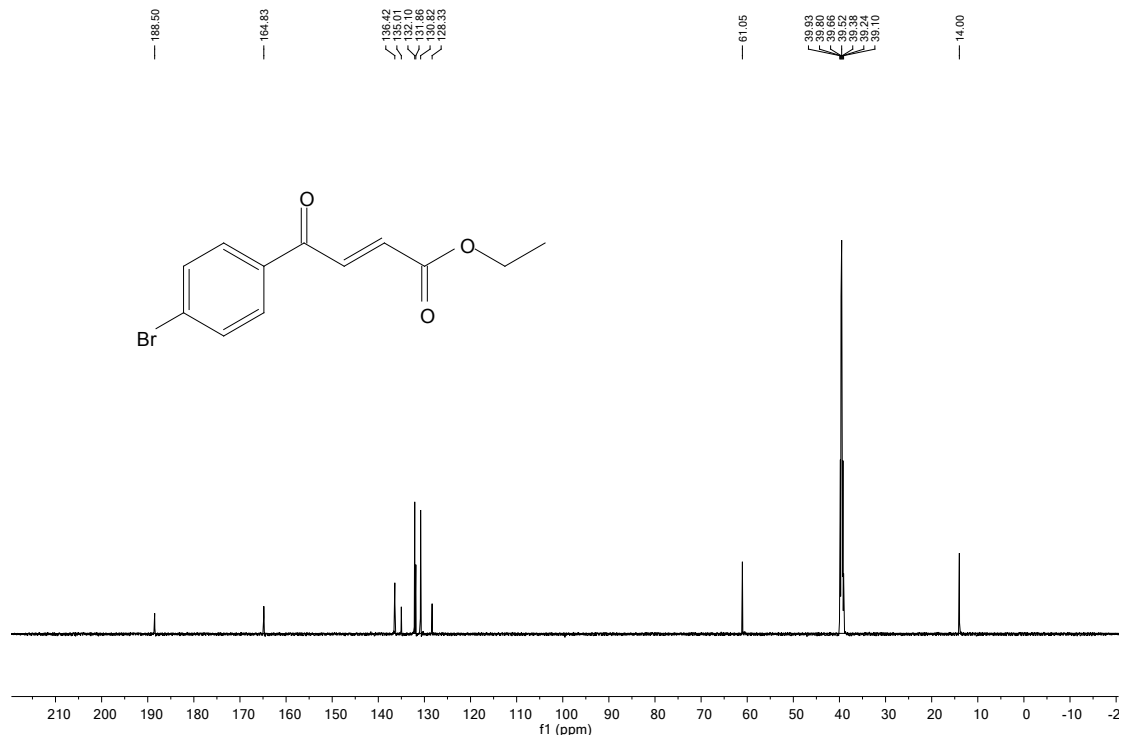
9o



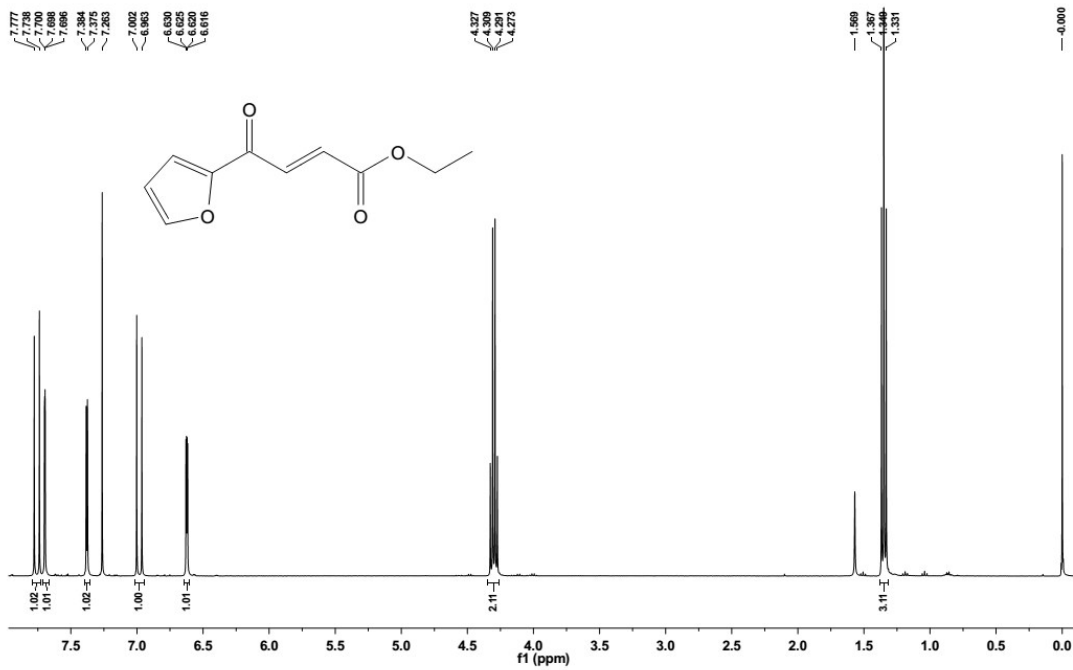
9p



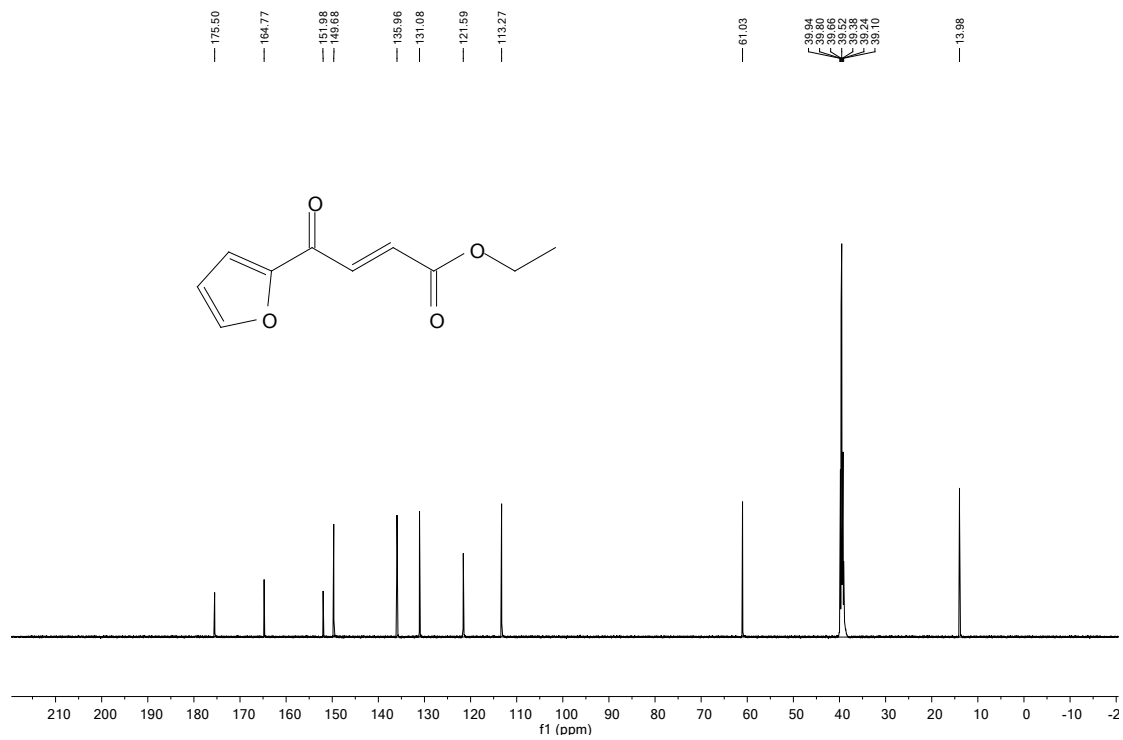
BRUKER AVANCE III 13C-NMR 9p IN DMSO-d6



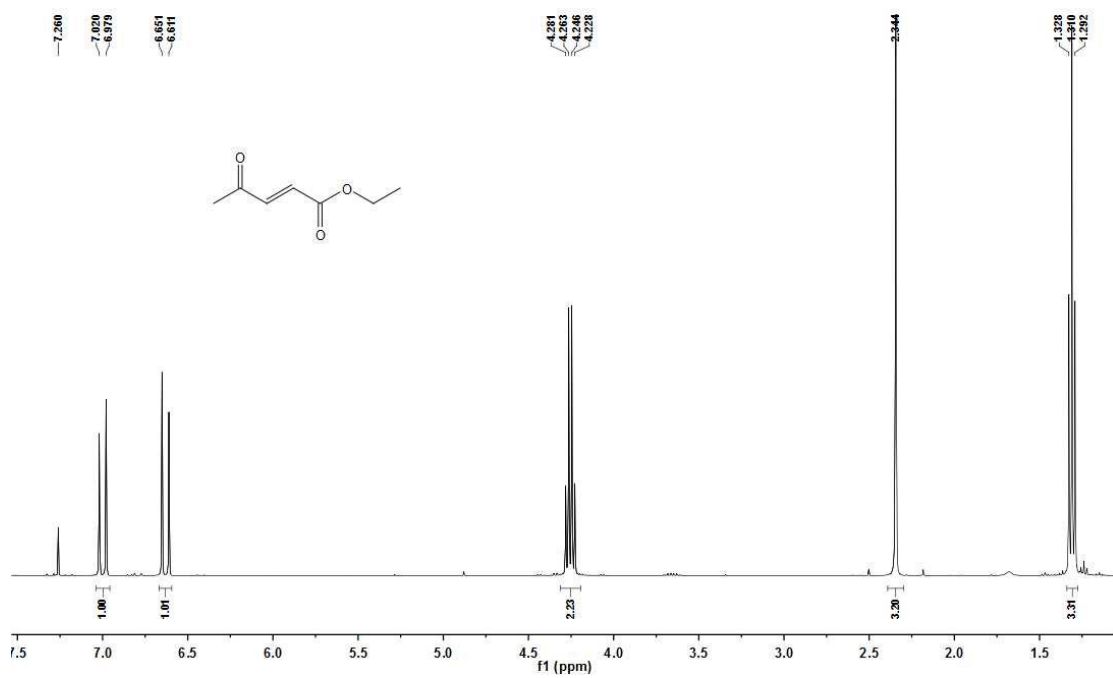
9q



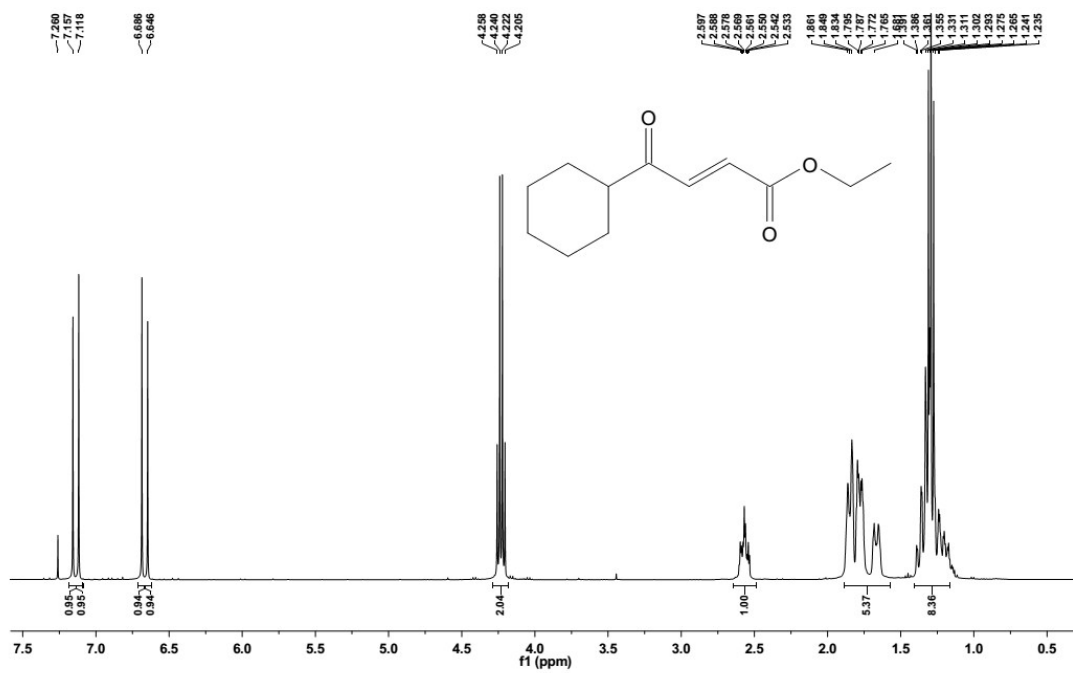
BRUKER AVANCE III 13C-NMR 9q IN DMSO-d6



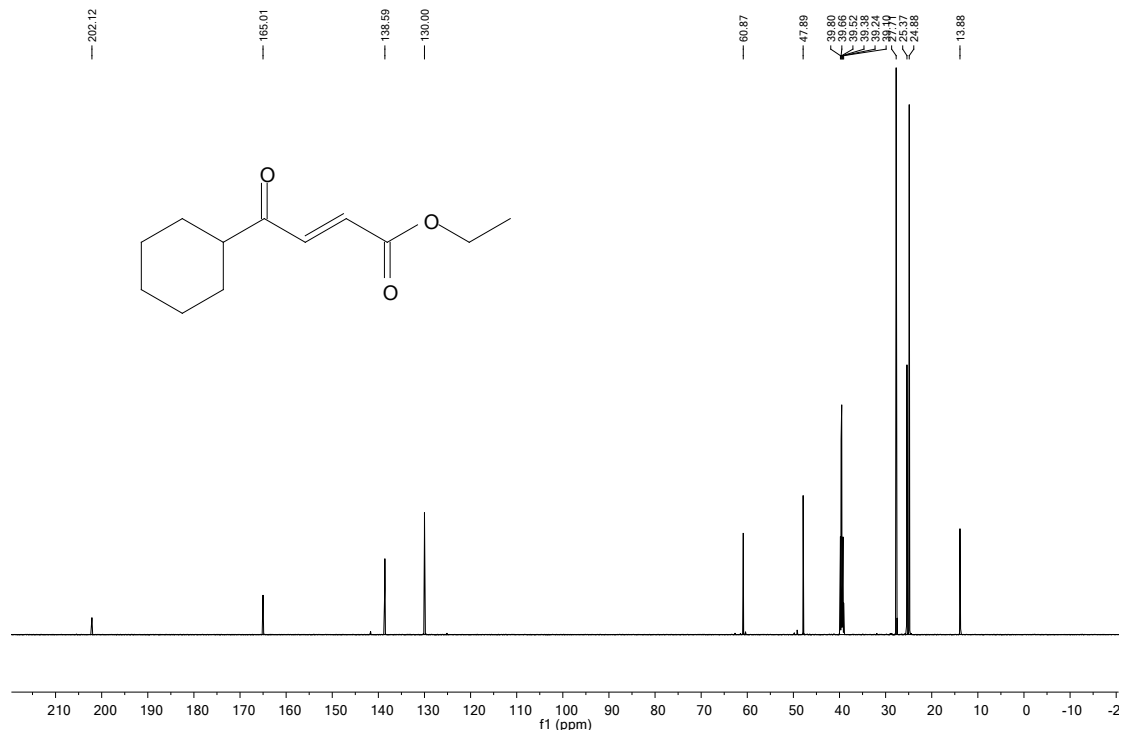
9r



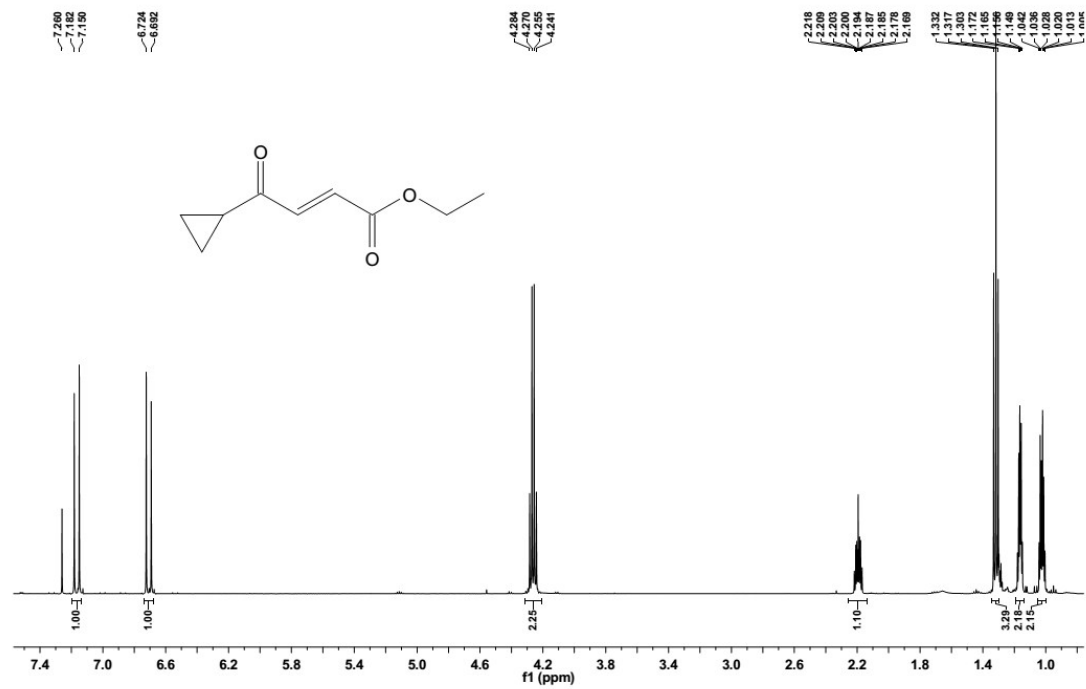
9s



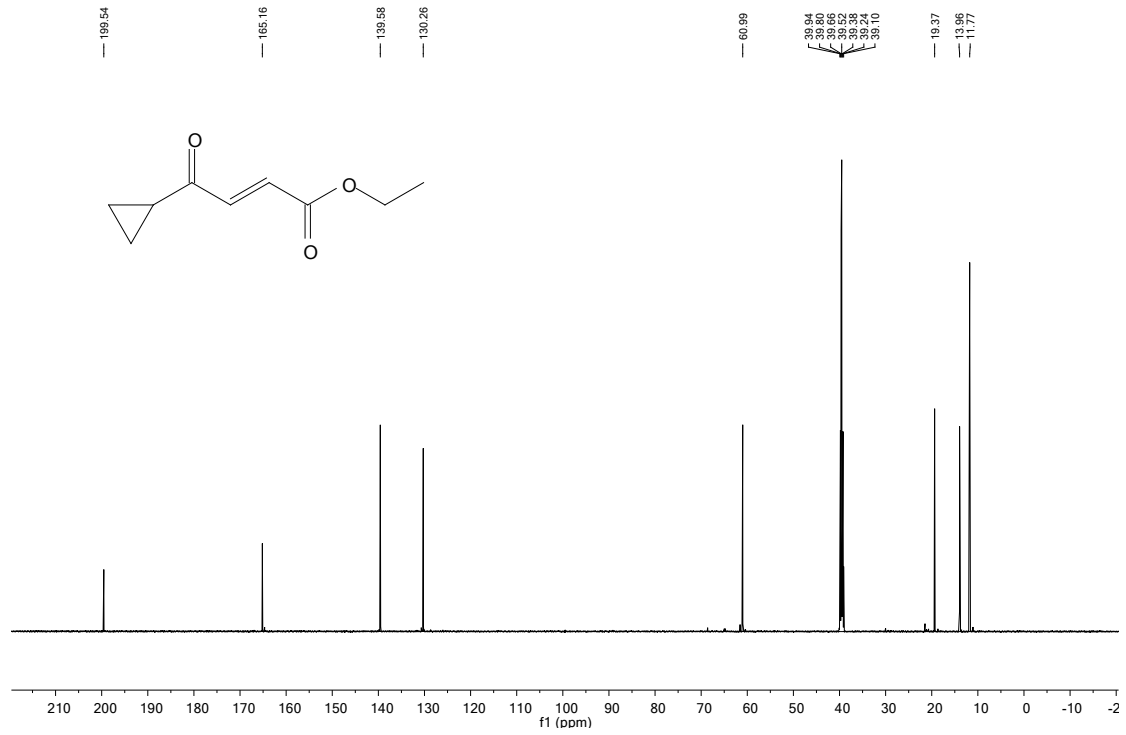
BRUKER AVANCE III 13C-NMR 9s IN DMSO-d6



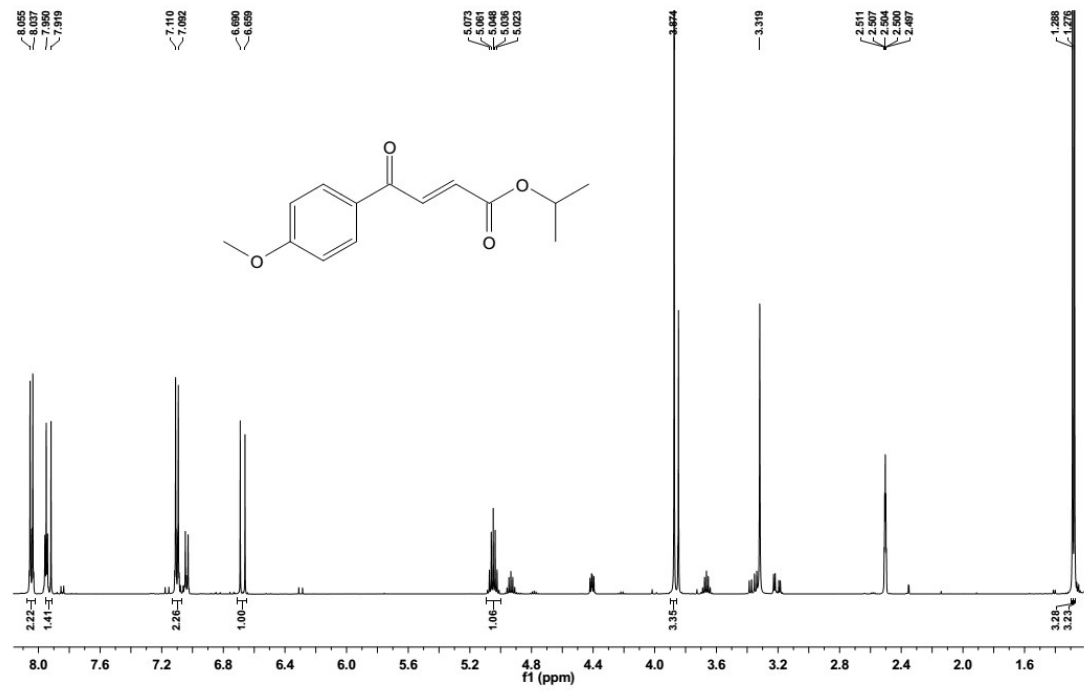
9t



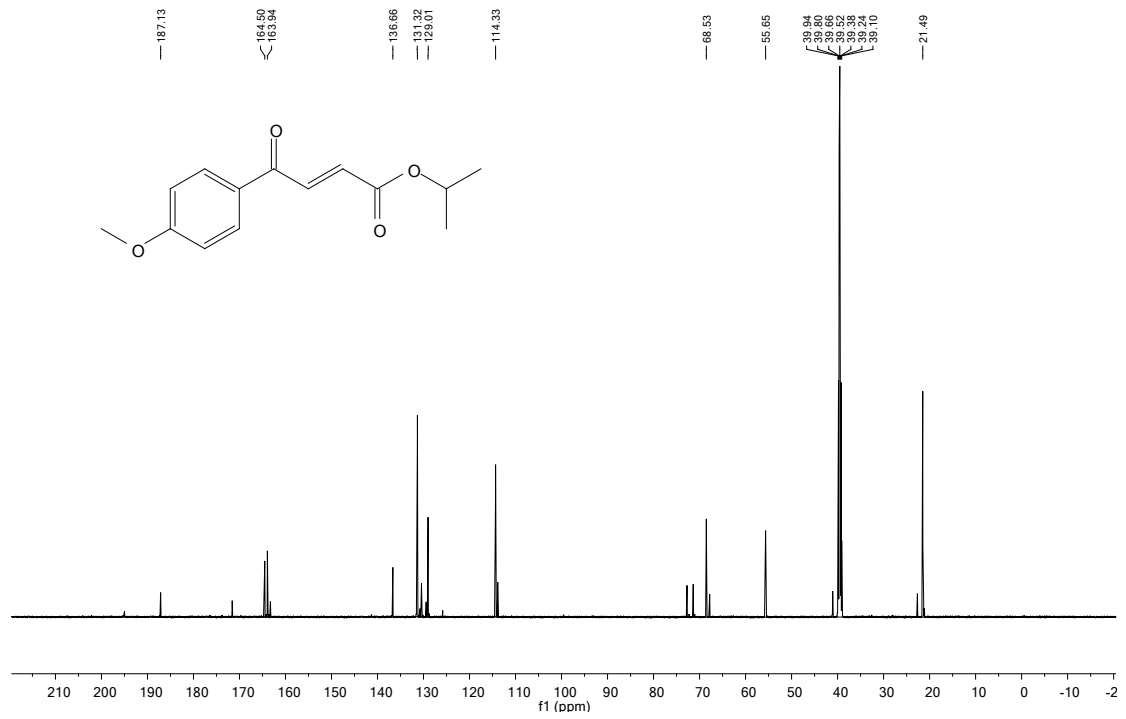
BRUKER AVANCE III 13C-NMR 9t IN DMSO-d6



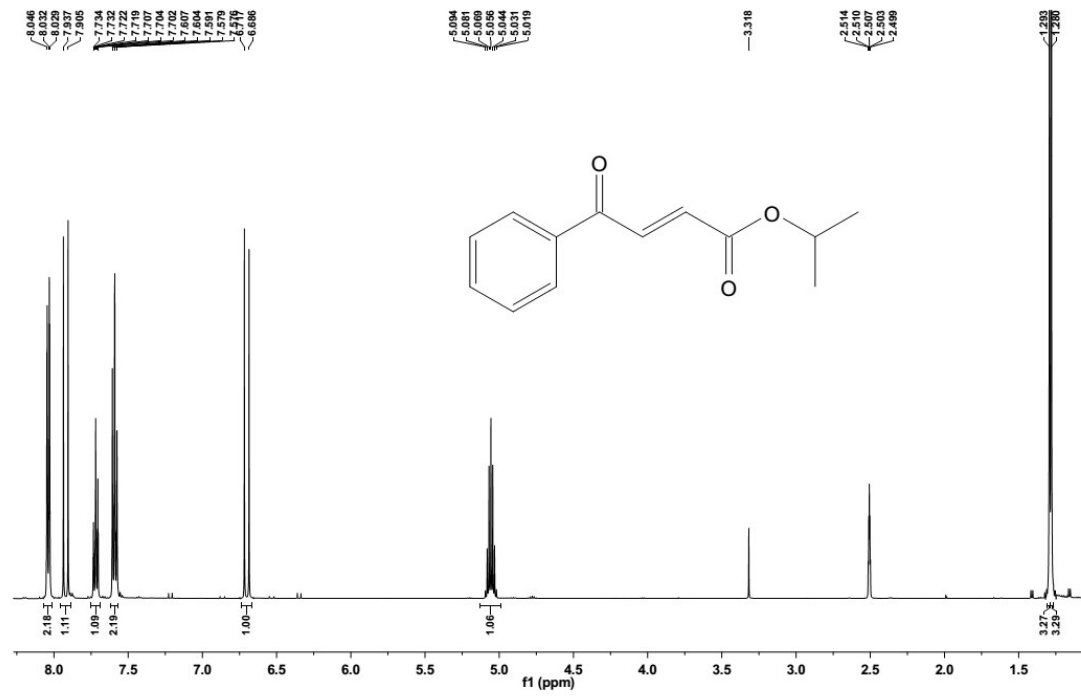
10a



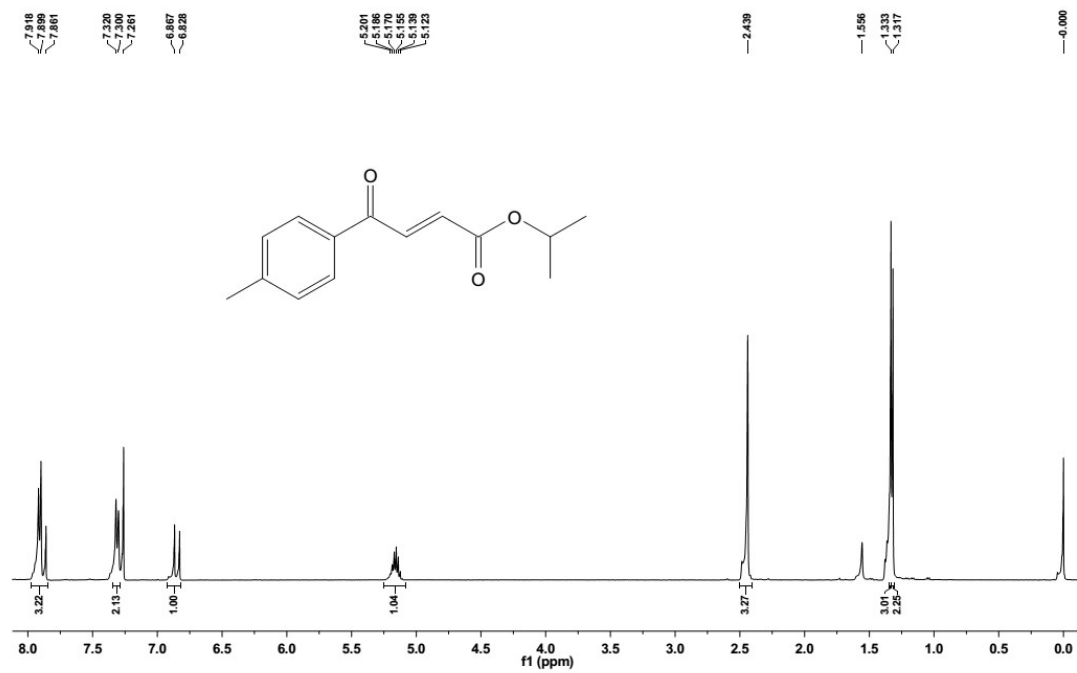
BRUKER AVANCE III 13C-NMR 10a IN DMSO-d6



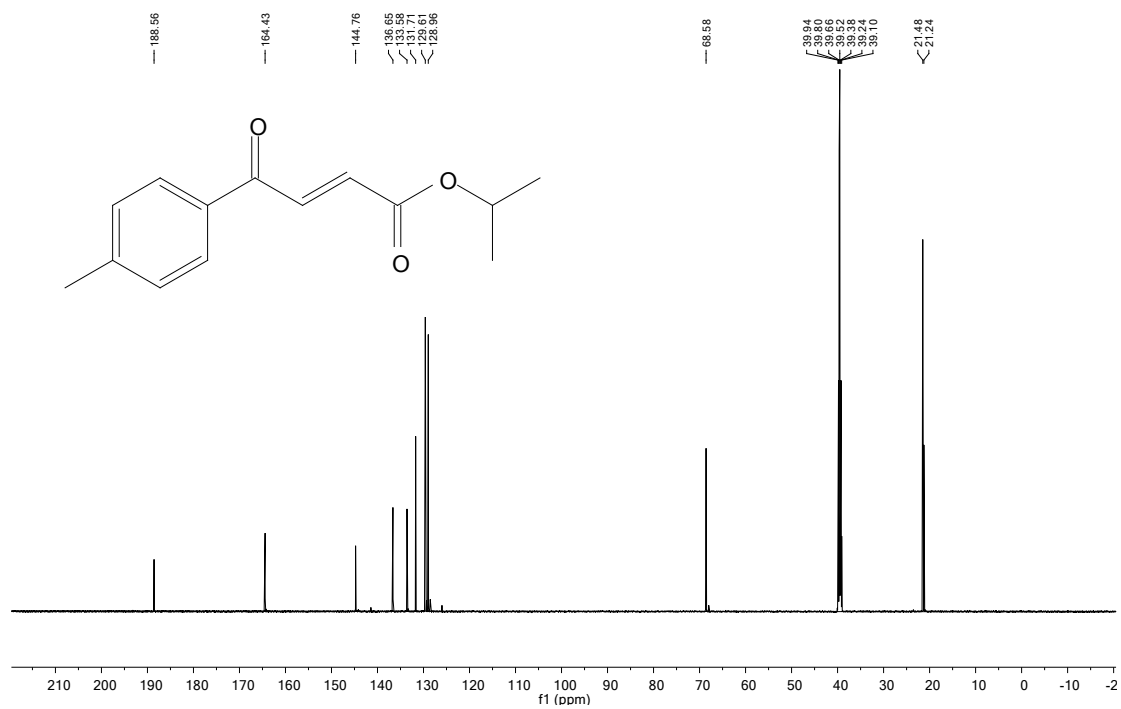
10b



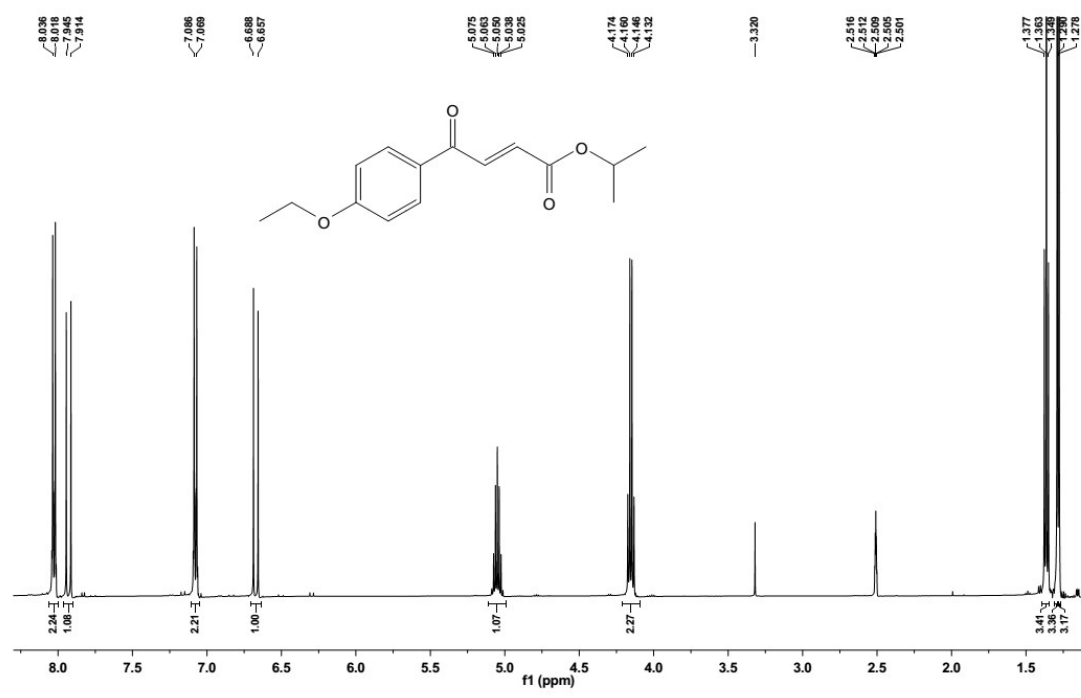
10c



BRUKER AVANCE III 13C-NMR 10c IN DMSO-d6



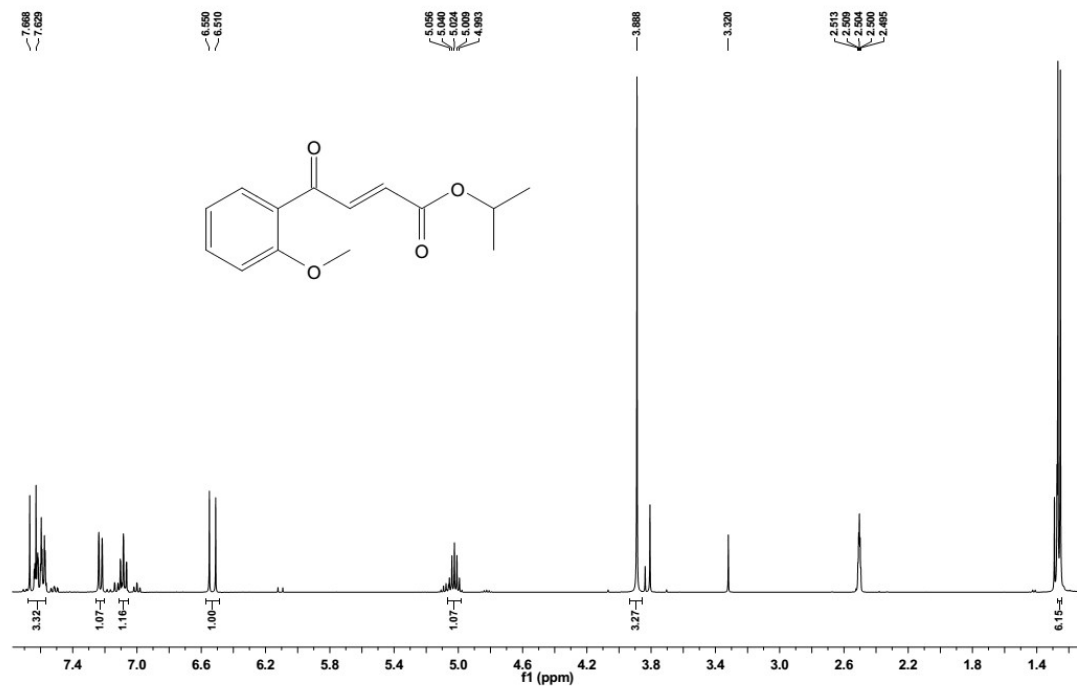
10d



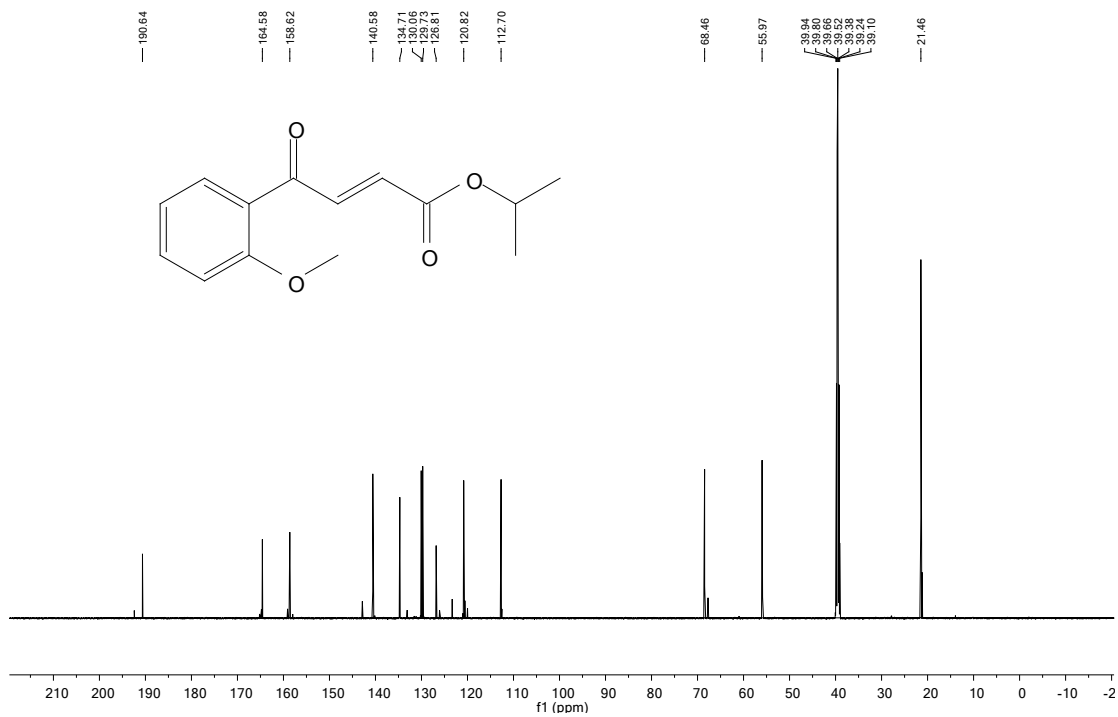
BRUKER AVANCE III 13C-NMR 10d IN DMSO-d6



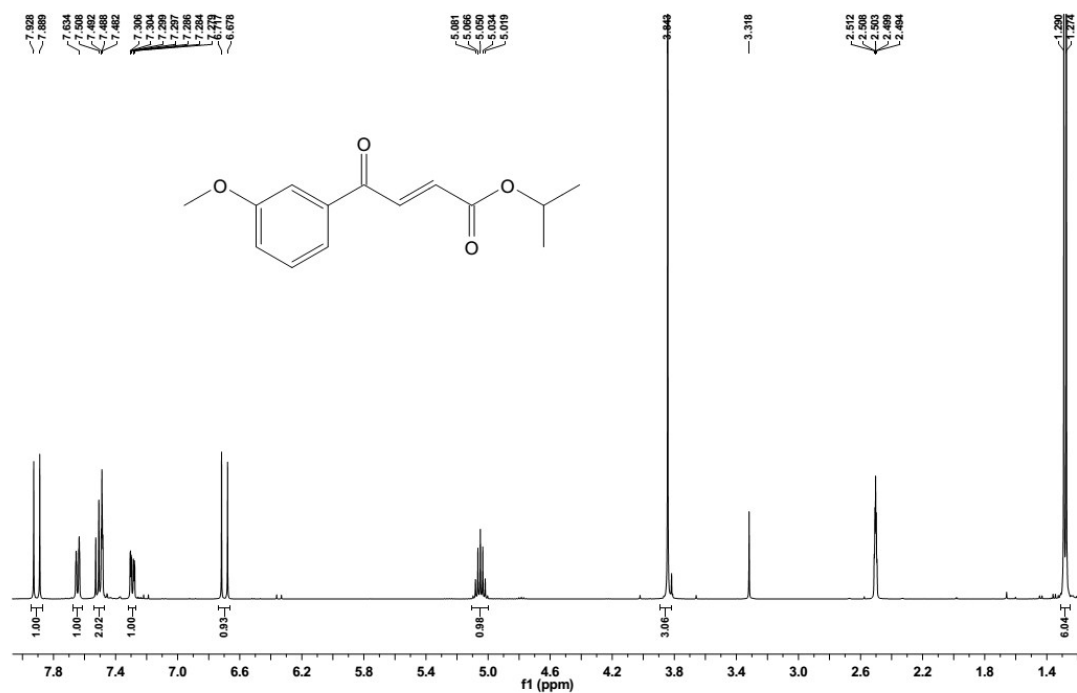
10e



BRUKER AVANCE III 13C-NMR 10e IN DMSO-d6



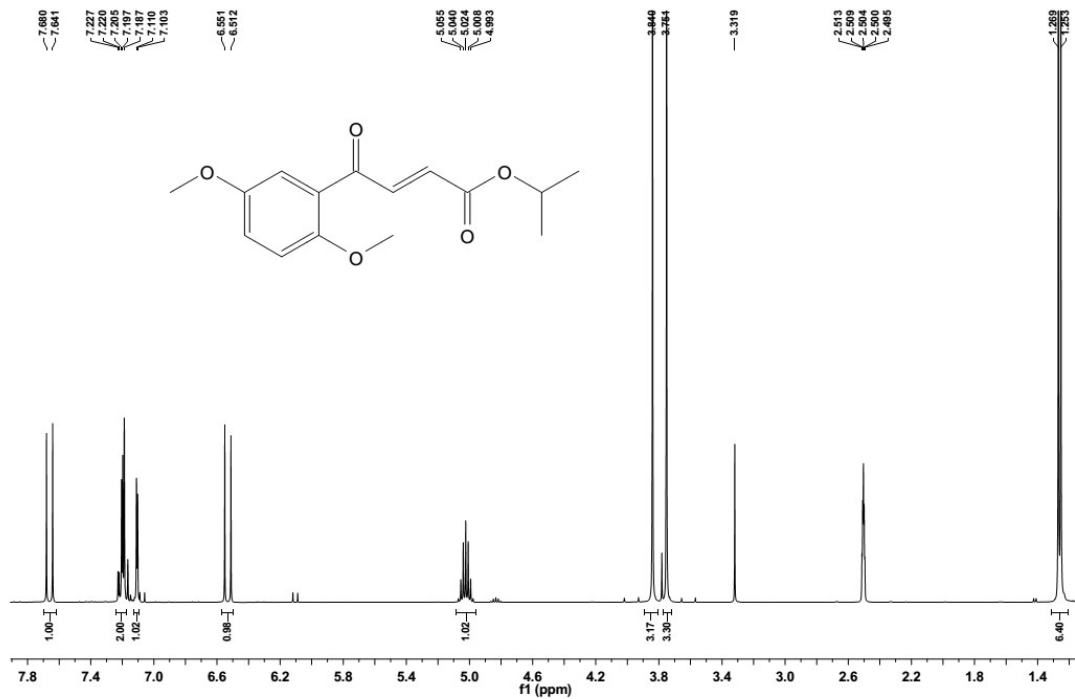
10f



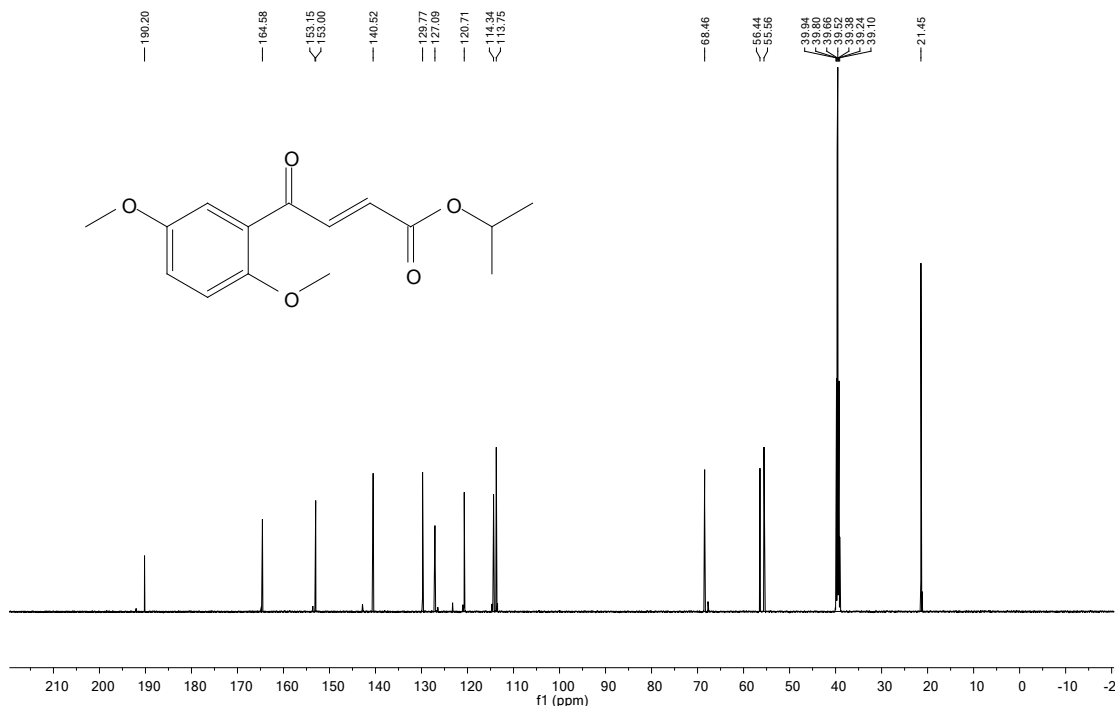
BRUKER AVANCE III 13C-NMR 10f IN DMSO-d6



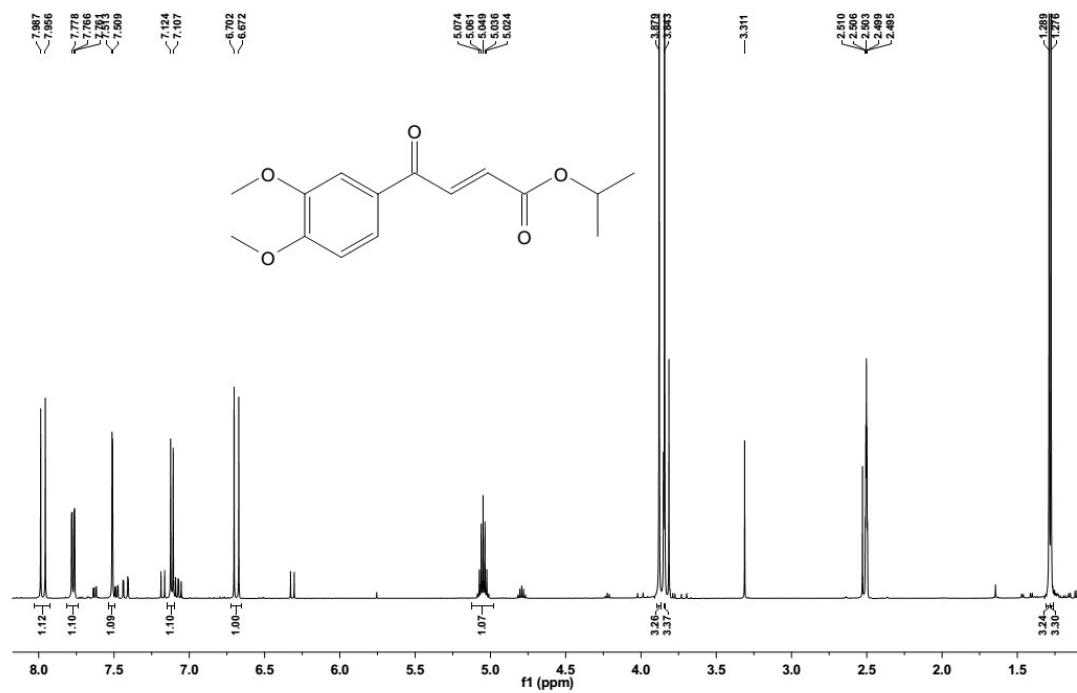
10g



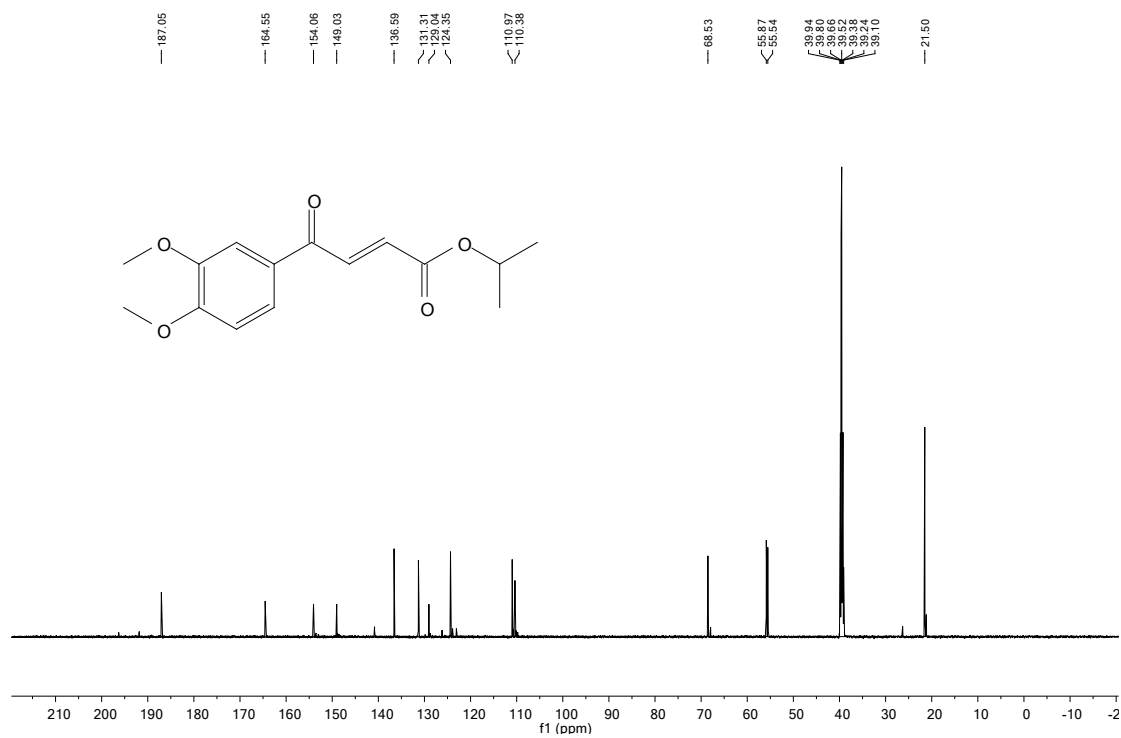
BRUKER AVANCE III 13C-NMR 10g IN DMSO-d6

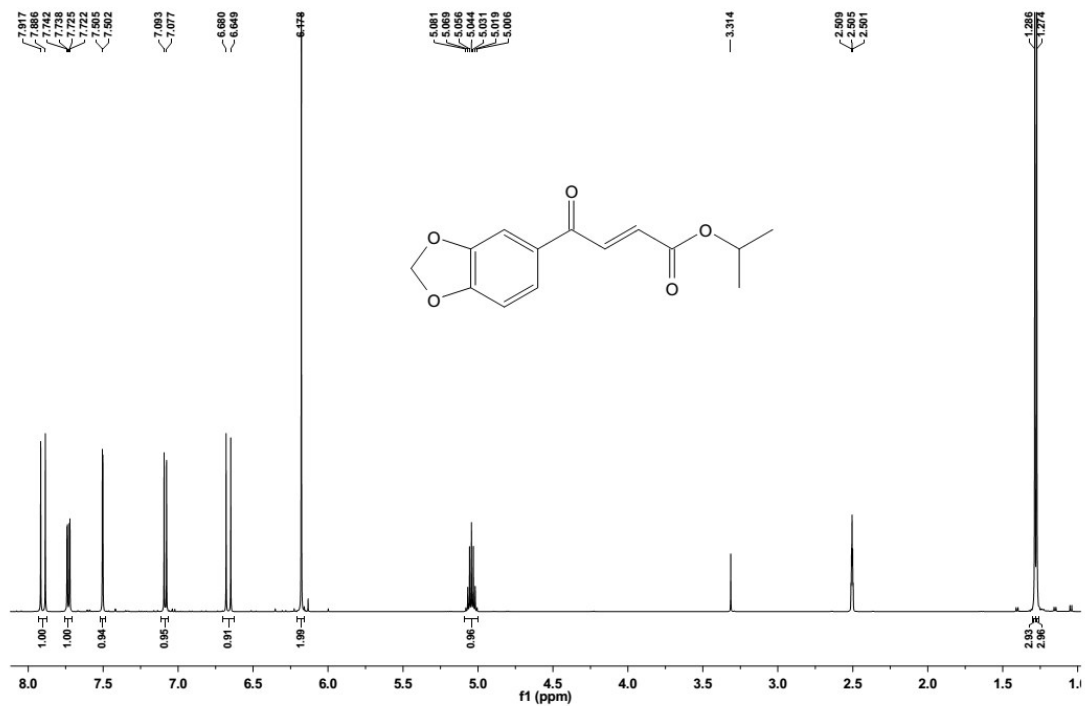


10h

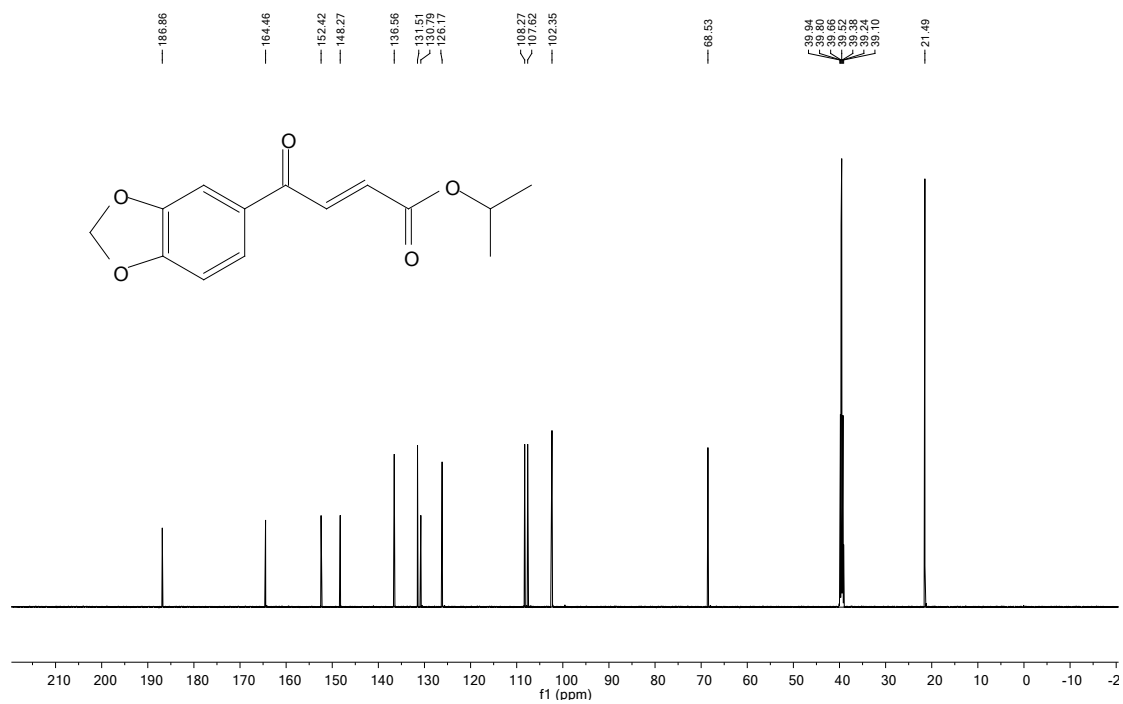


BRUKER AVANCE III 13C-NMR 10h IN DMSO-d6

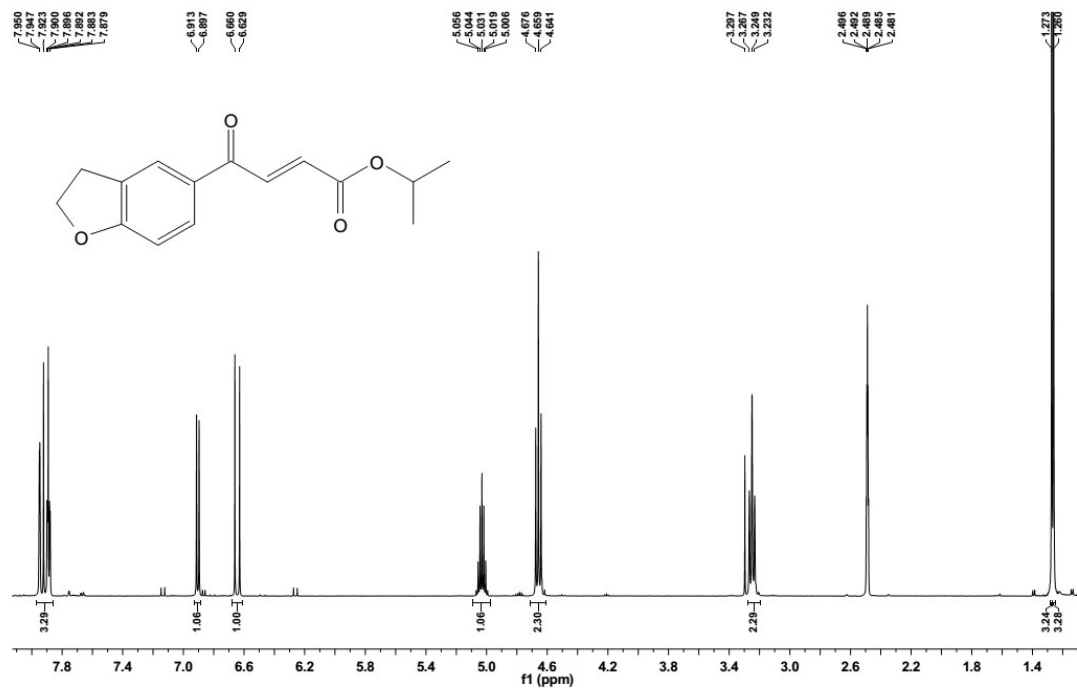




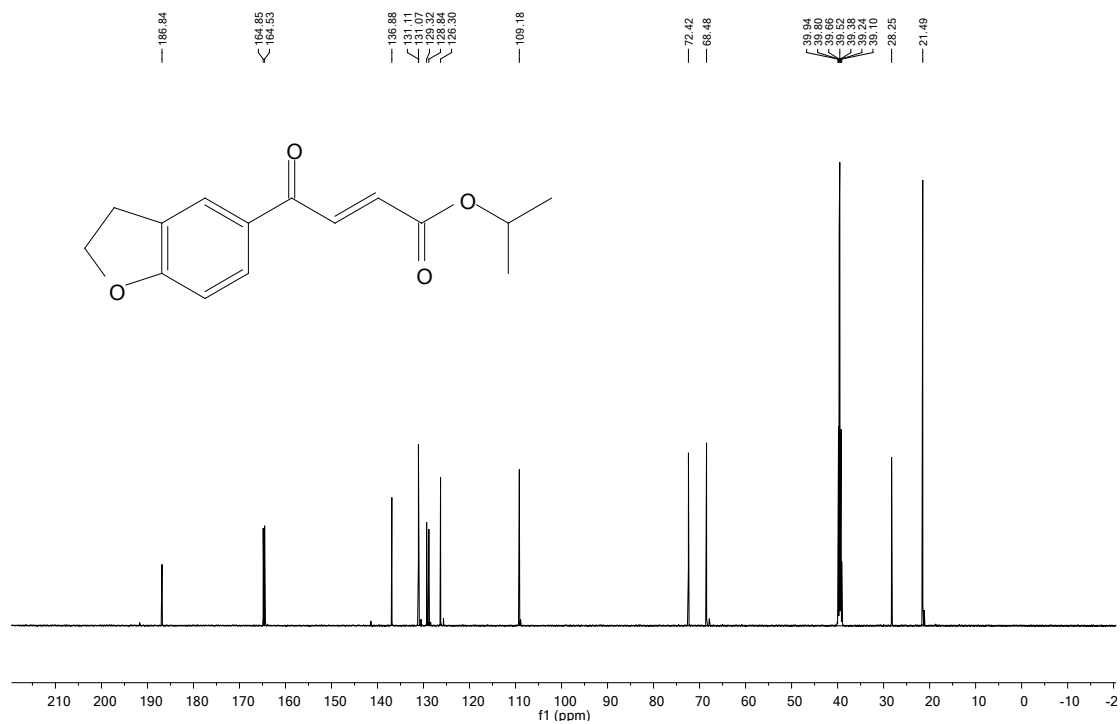
BRUKER AVANCE III 13C-NMR 10i IN DMSO-d6



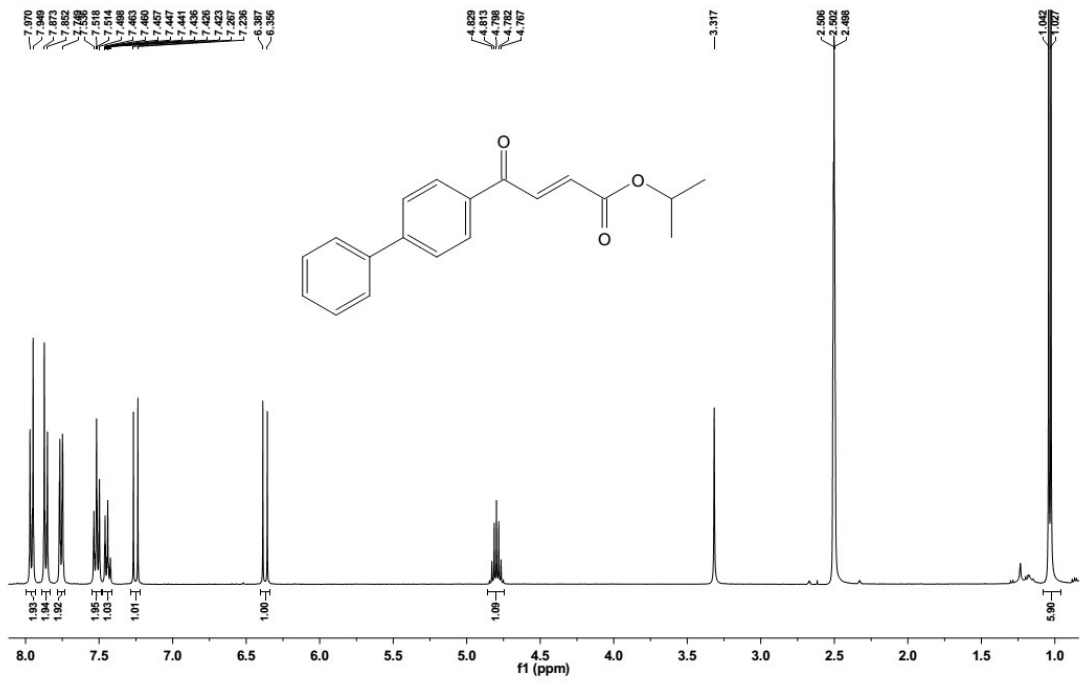
10j



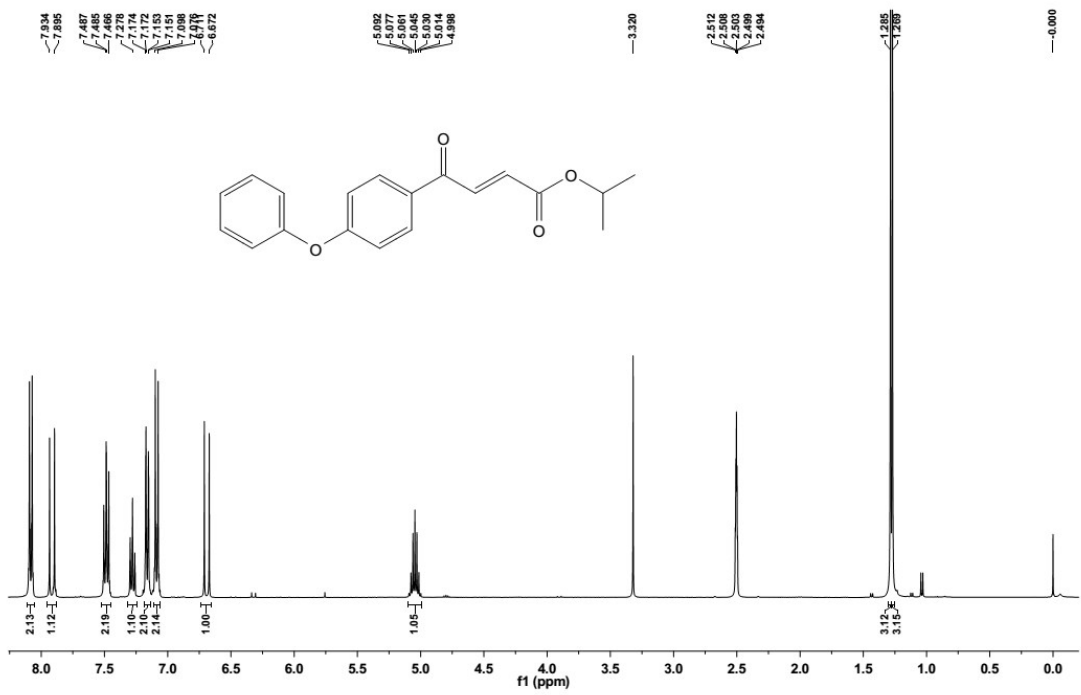
BRUKER AVANCE III 13C-NMR 10j IN DMSO-d₆



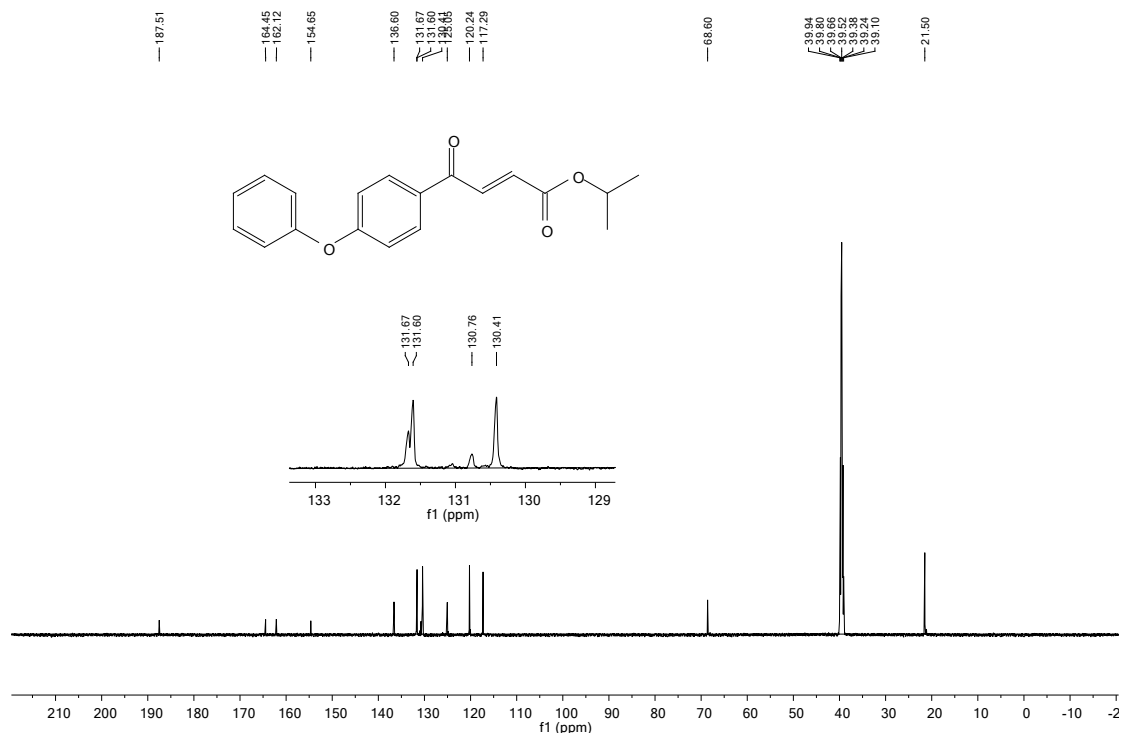
10k



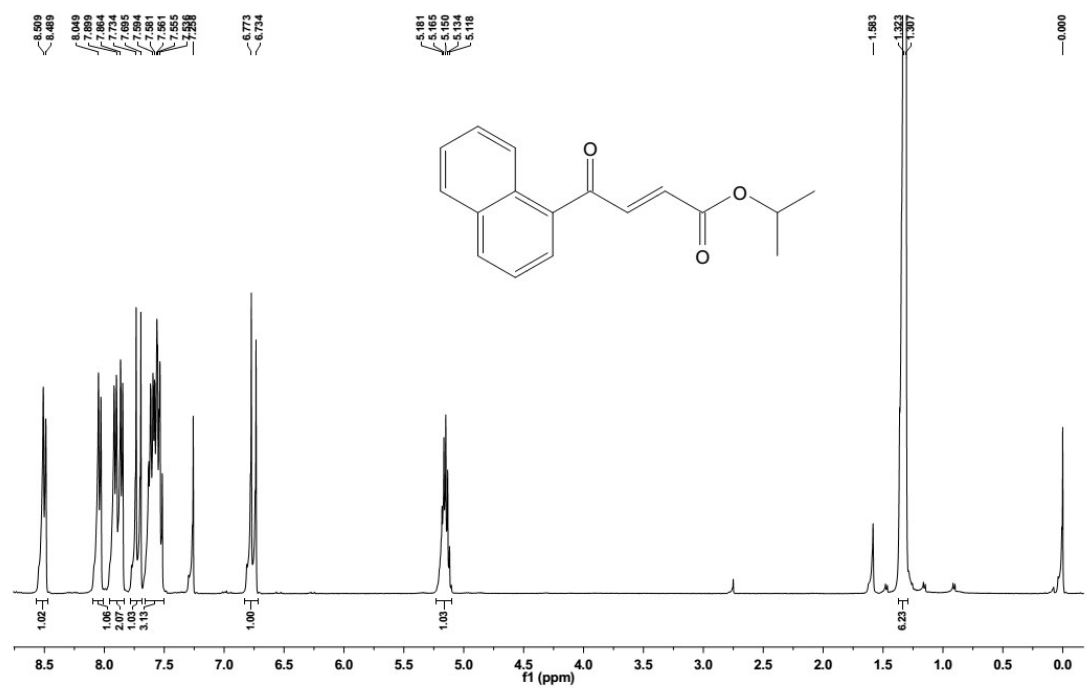
101



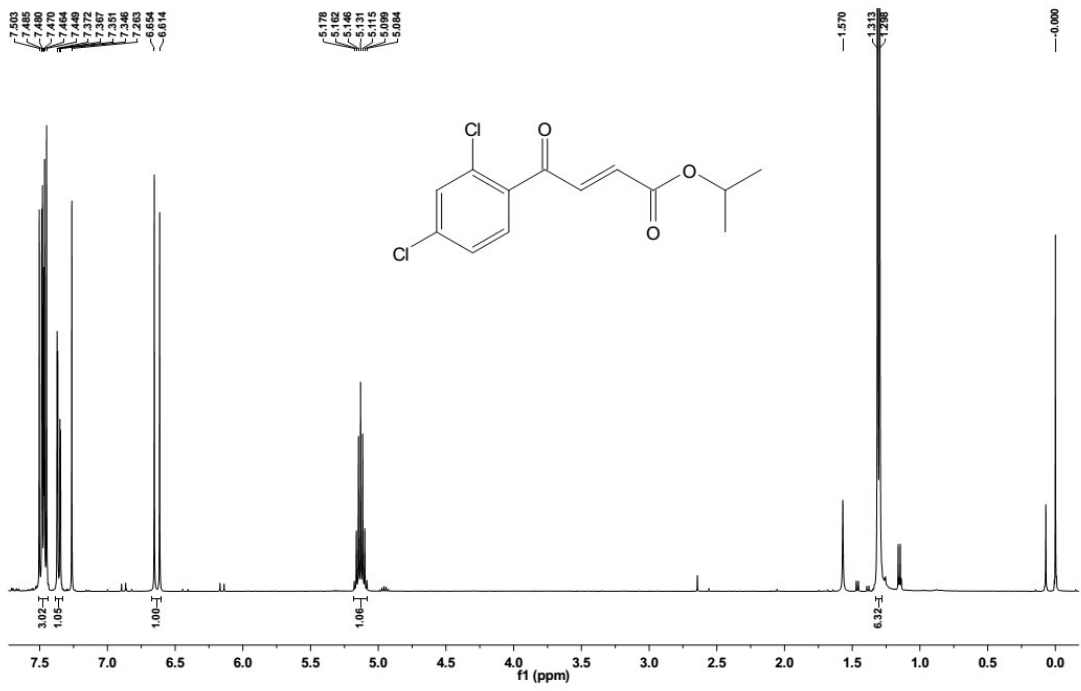
BRUKER AVANCE III 13C-NMR 101 IN DMSO-d6



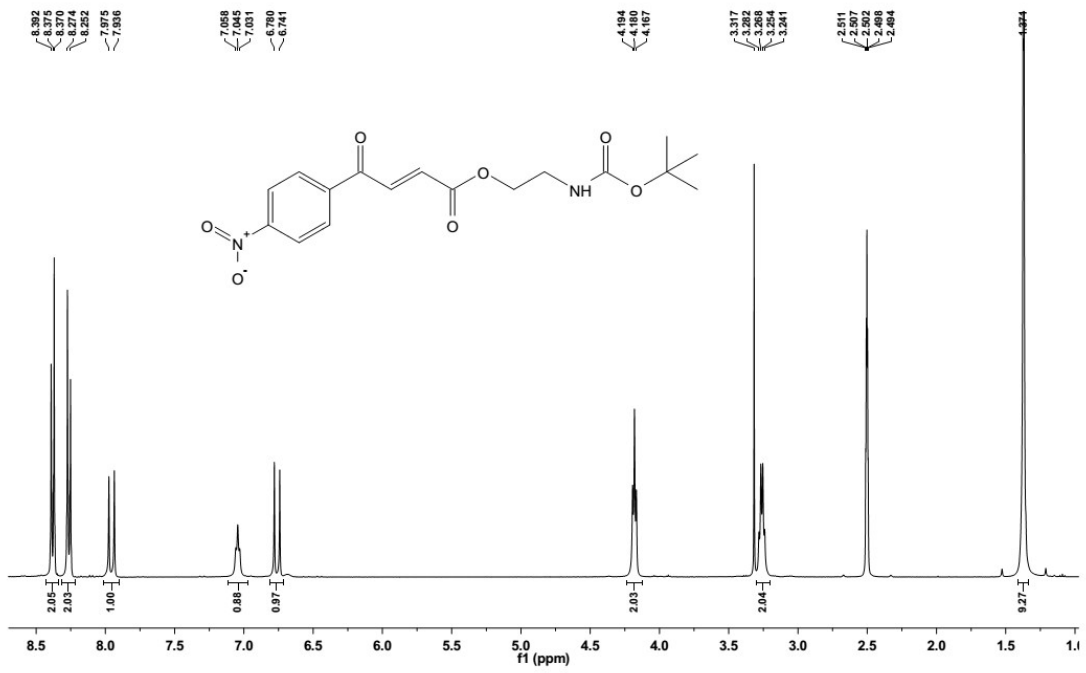
10m



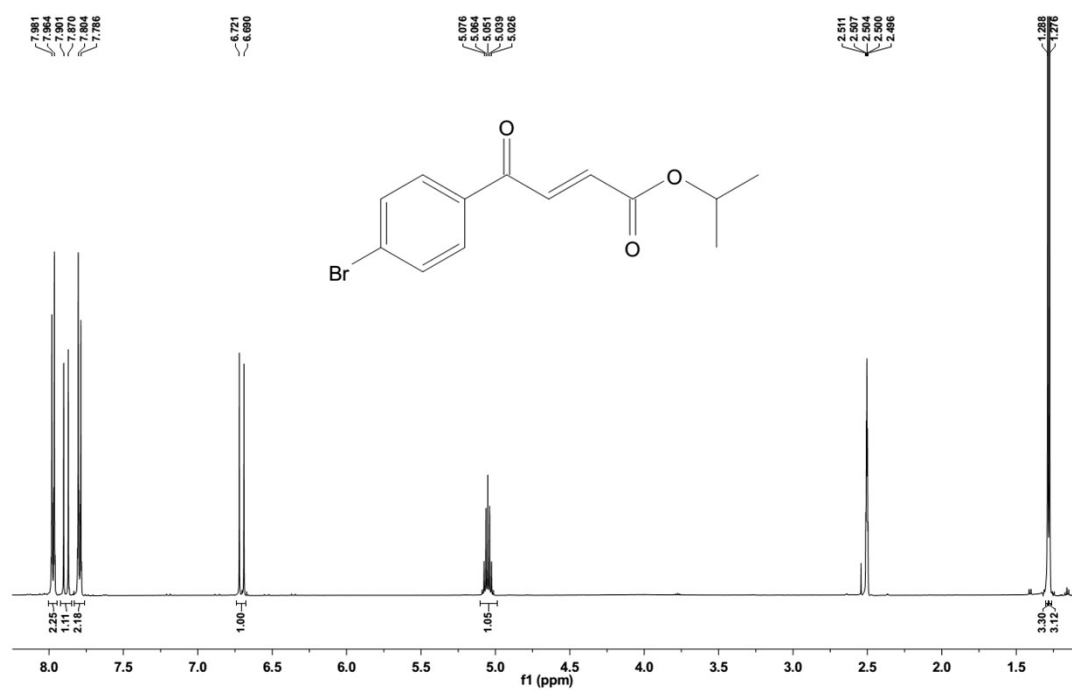
10n



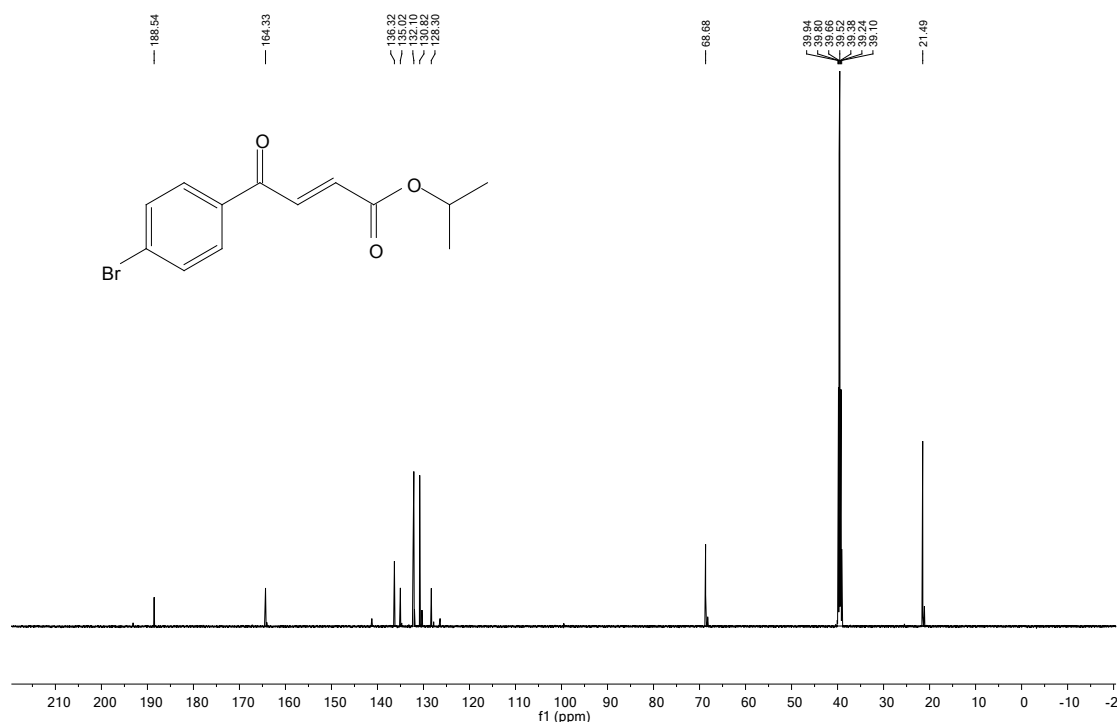
10o



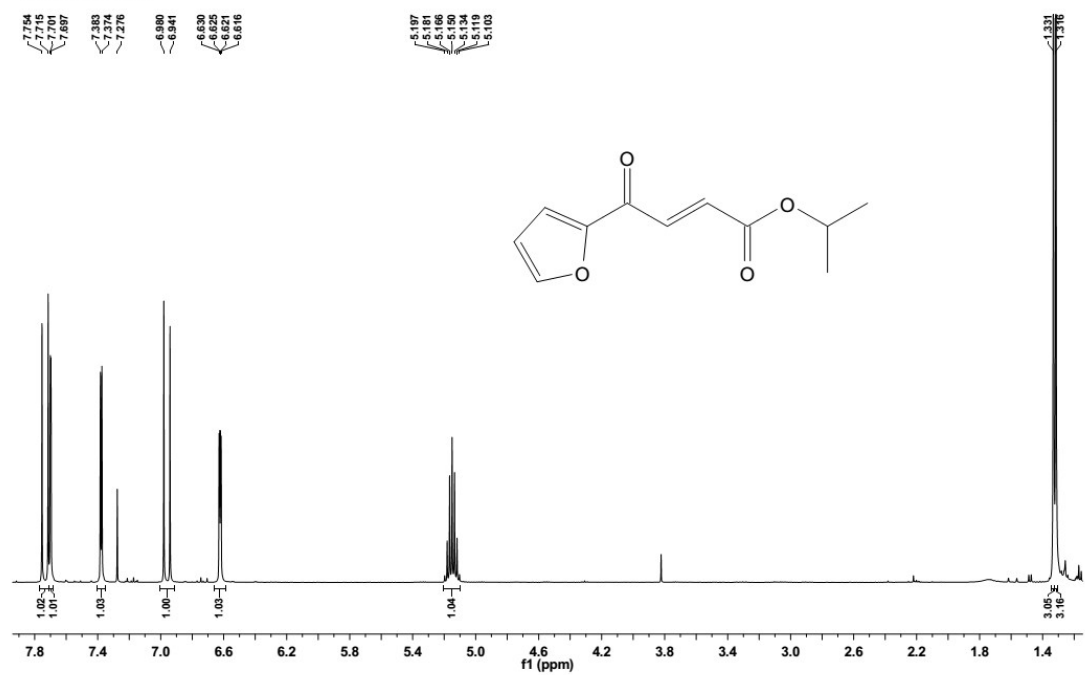
10p



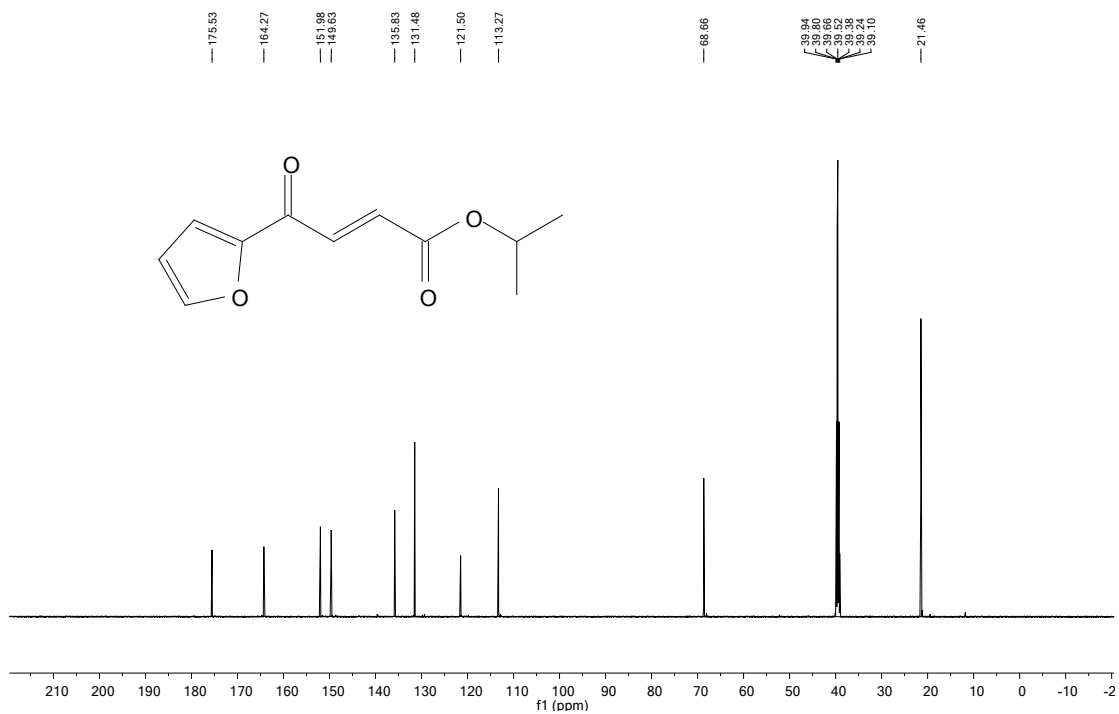
BRUKER AVANCE III 13C-NMR 10p IN DMSO-d6



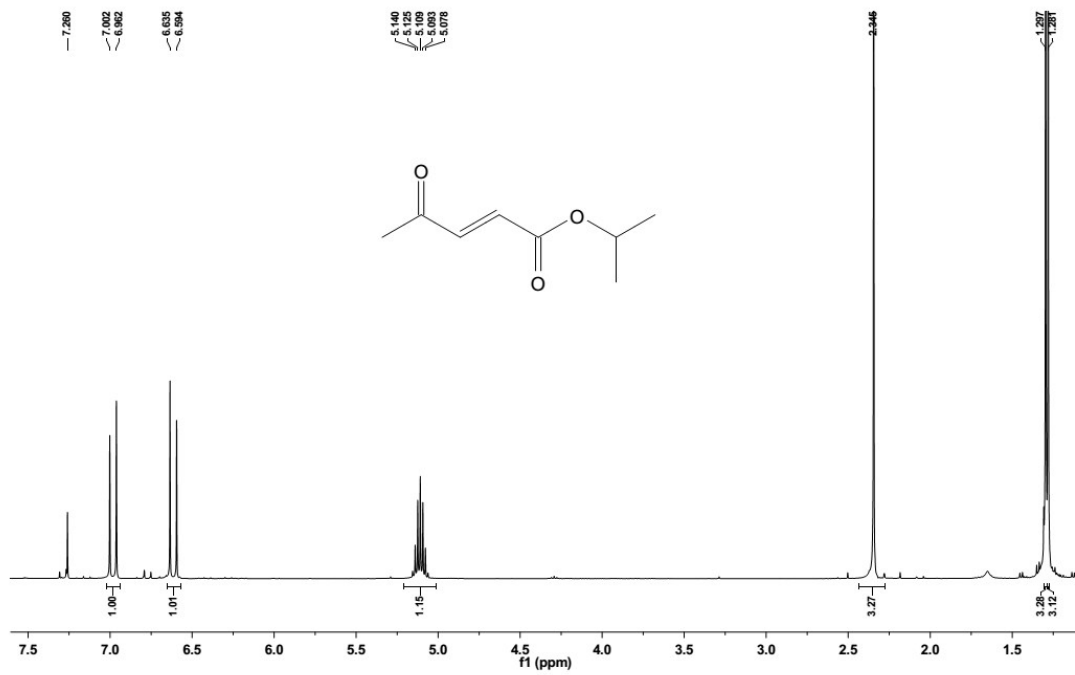
10q



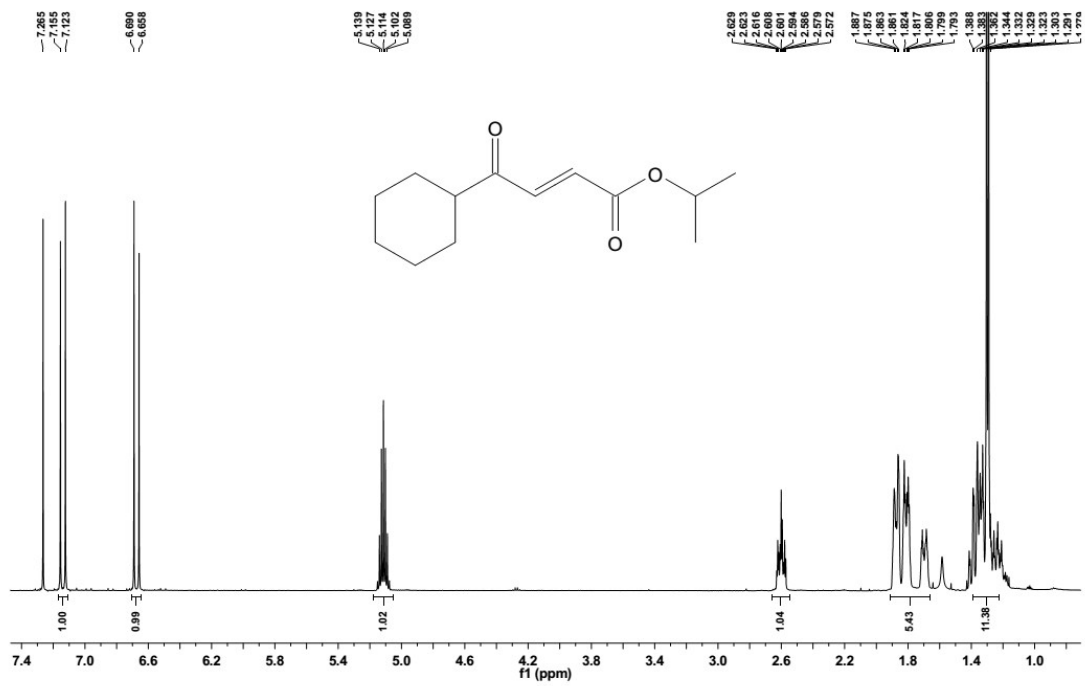
BRUKER AVANCE III 13C-NMR 10q IN DMSO-d6



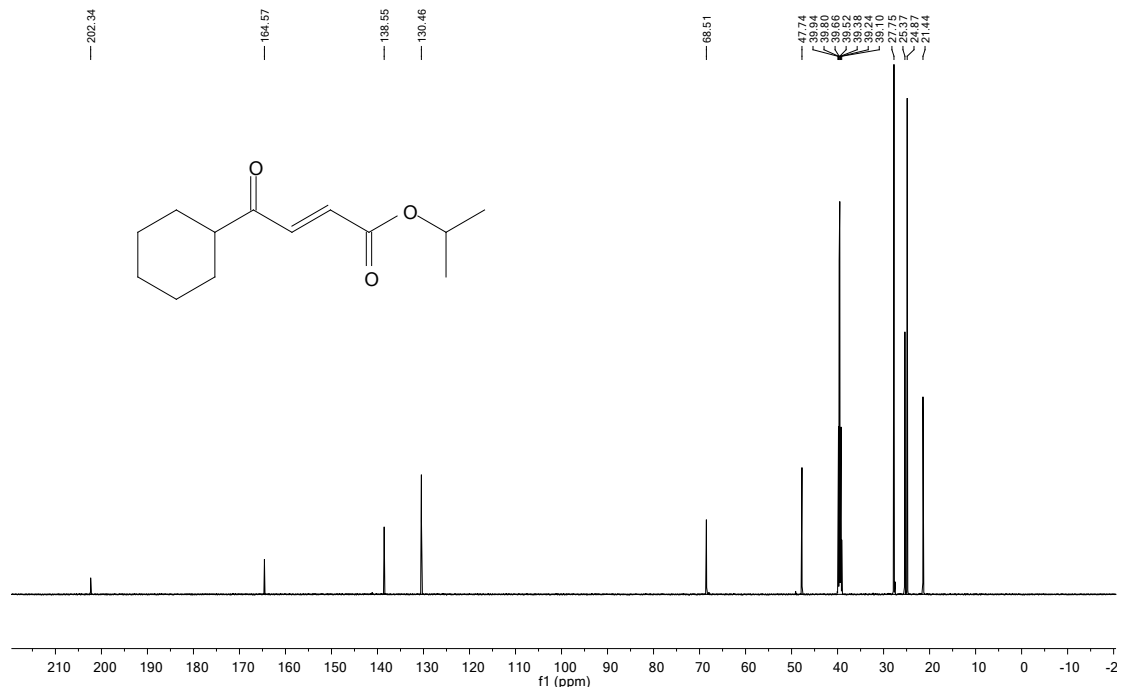
10r



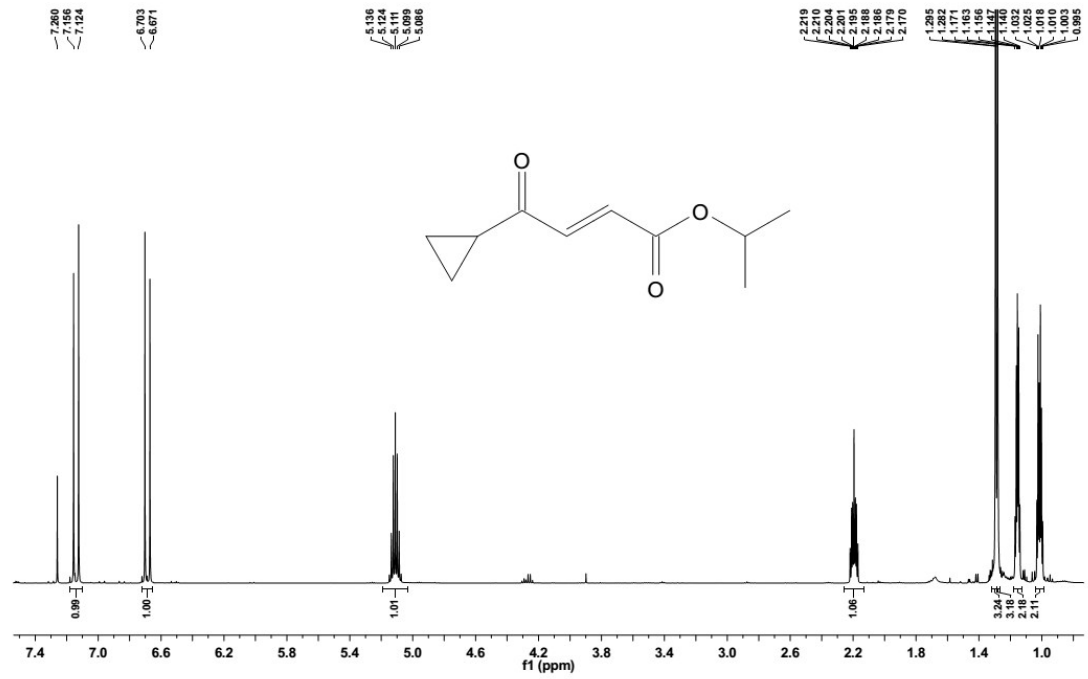
10s



BRUKER AVANCE III 13C-NMR 10s IN DMSO-d6



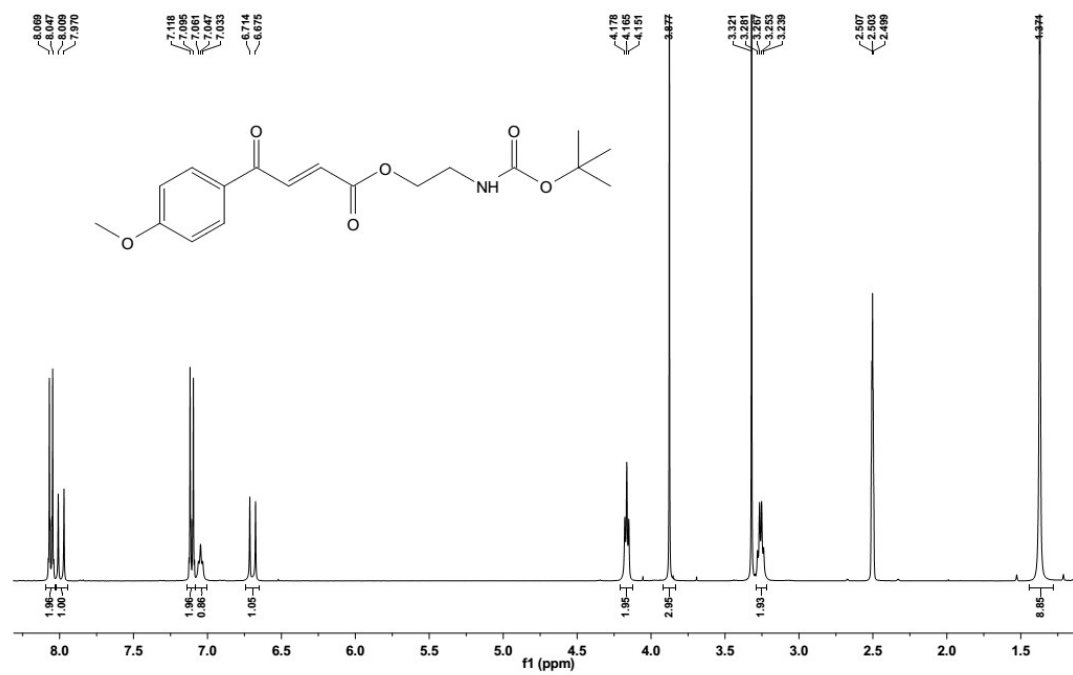
10t



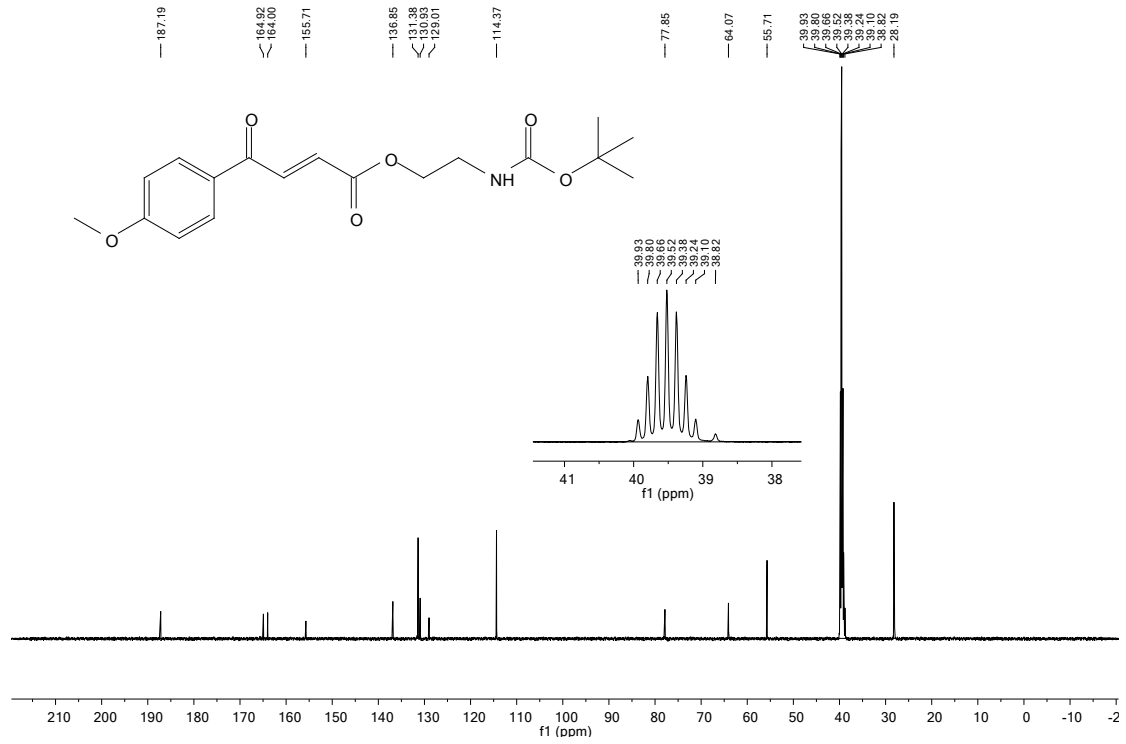
BRUKER AVANCE III 13C-NMR 10t IN DMSO-d6



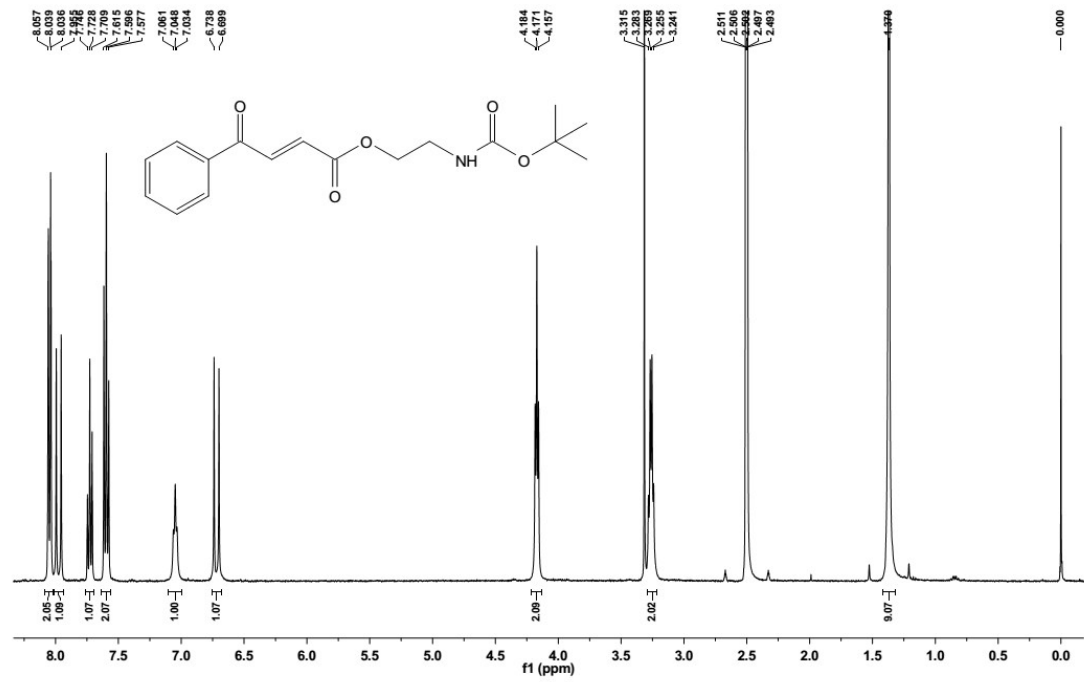
11a



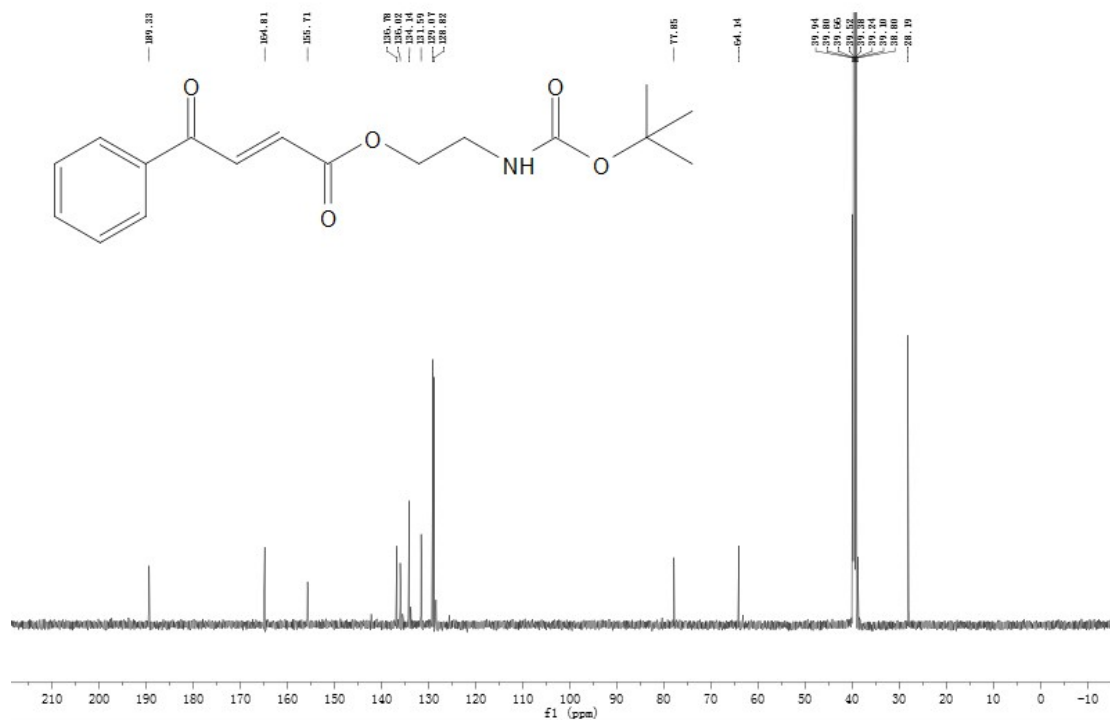
BRUKER AVANCE III 13C-NMR 11a IN DMSO-d6



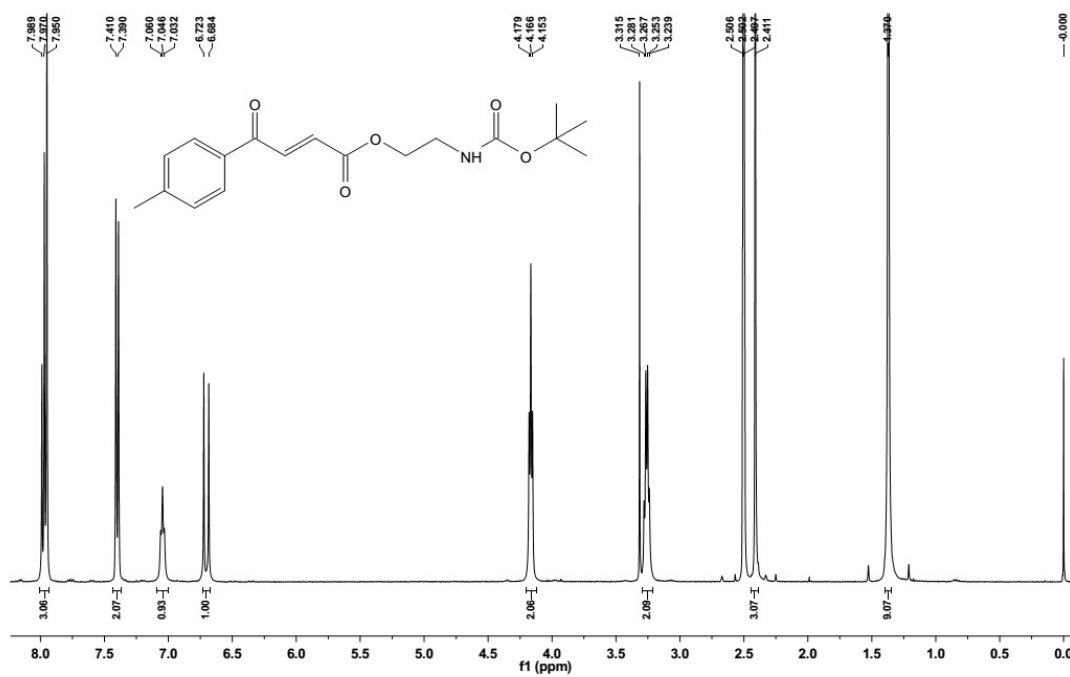
11b



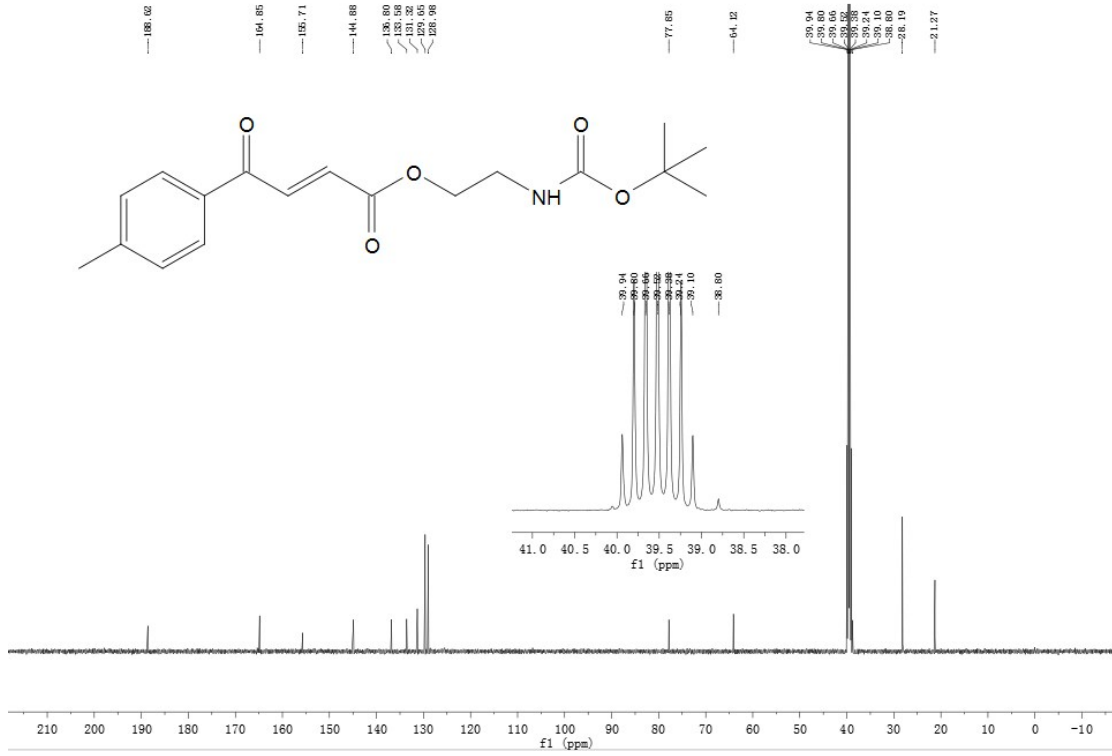
BRUKER AVANCE III 13C-NMR 11b IN DMSO-d6



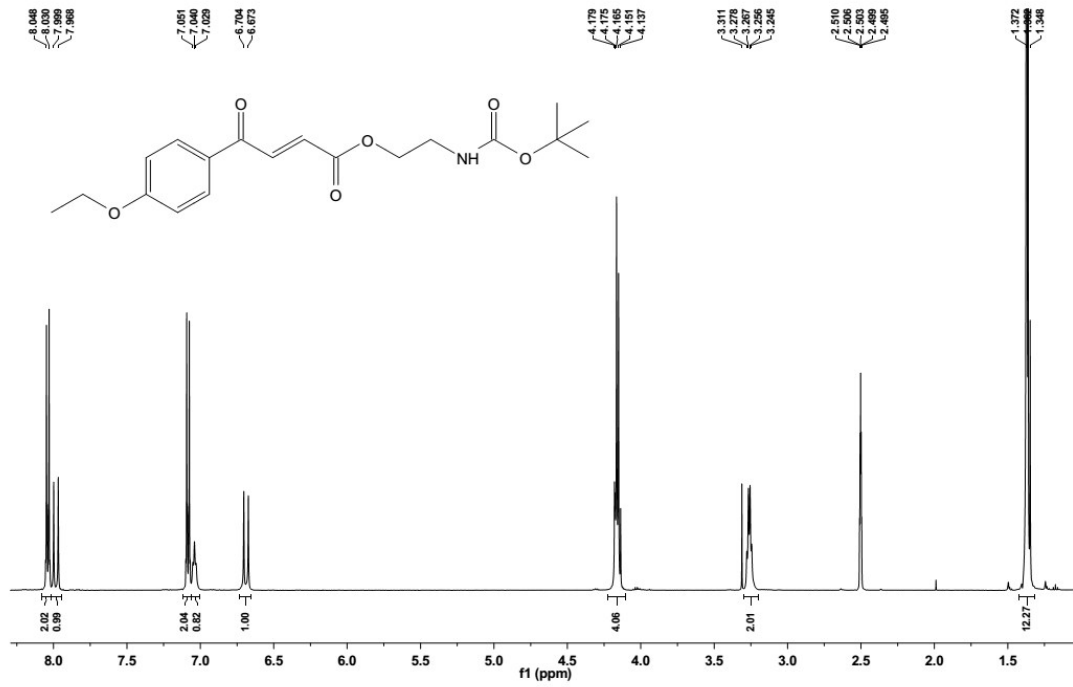
11c



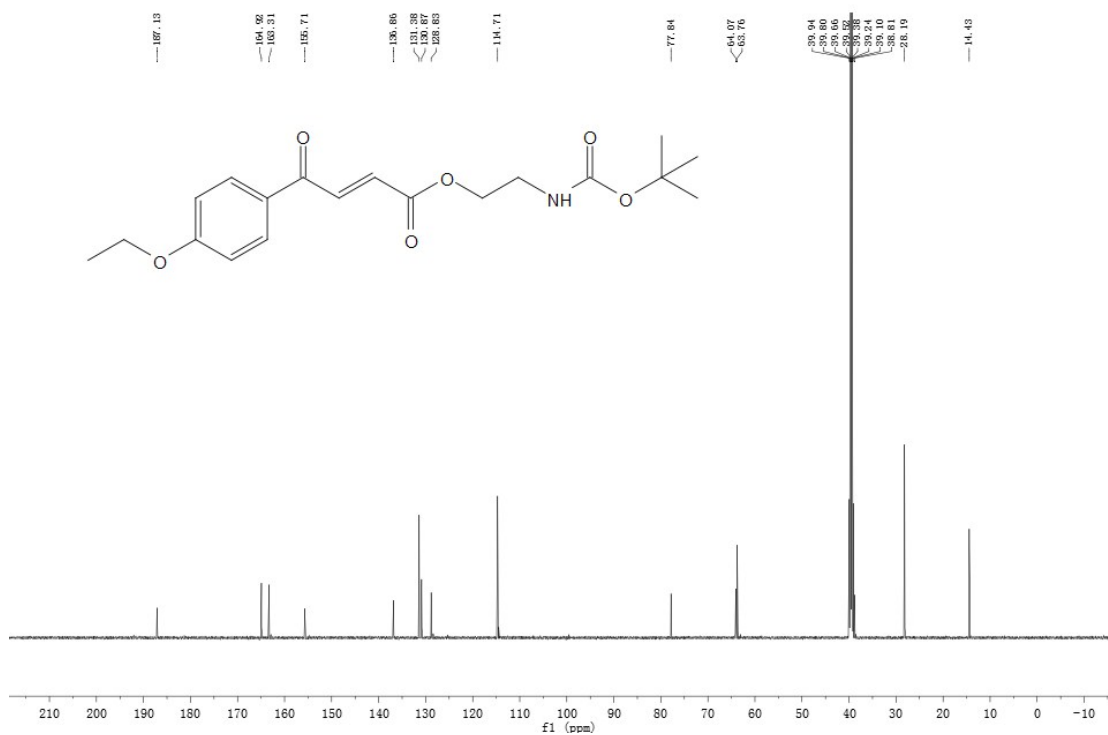
BRUKER AVANCE III 13C-NMR 11c IN DMSO-d6



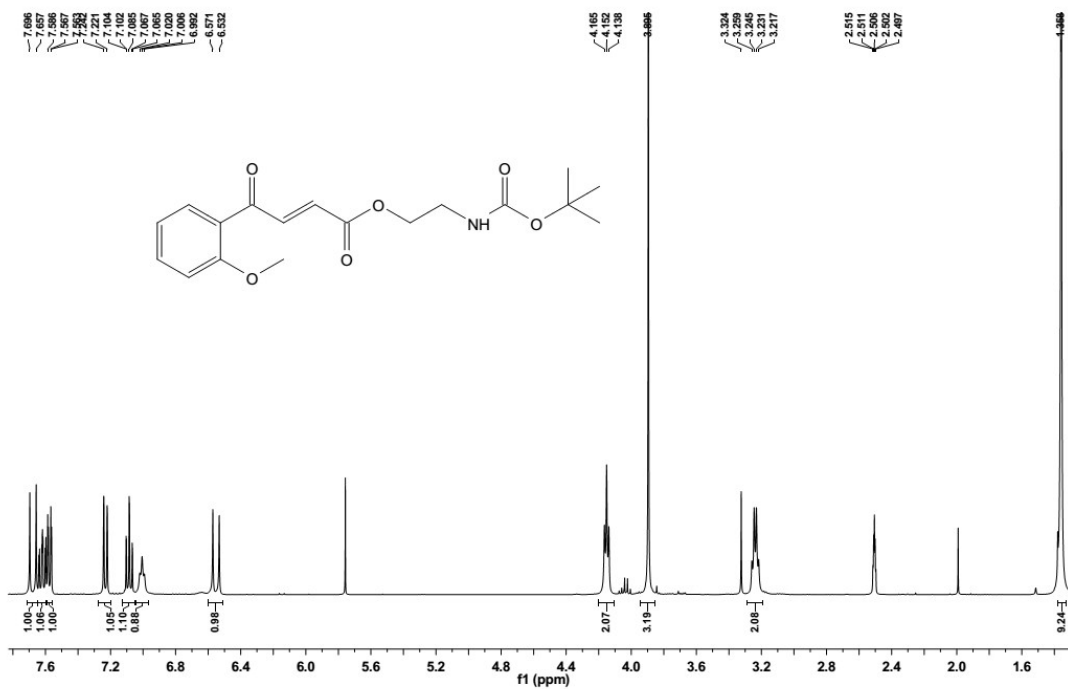
11d



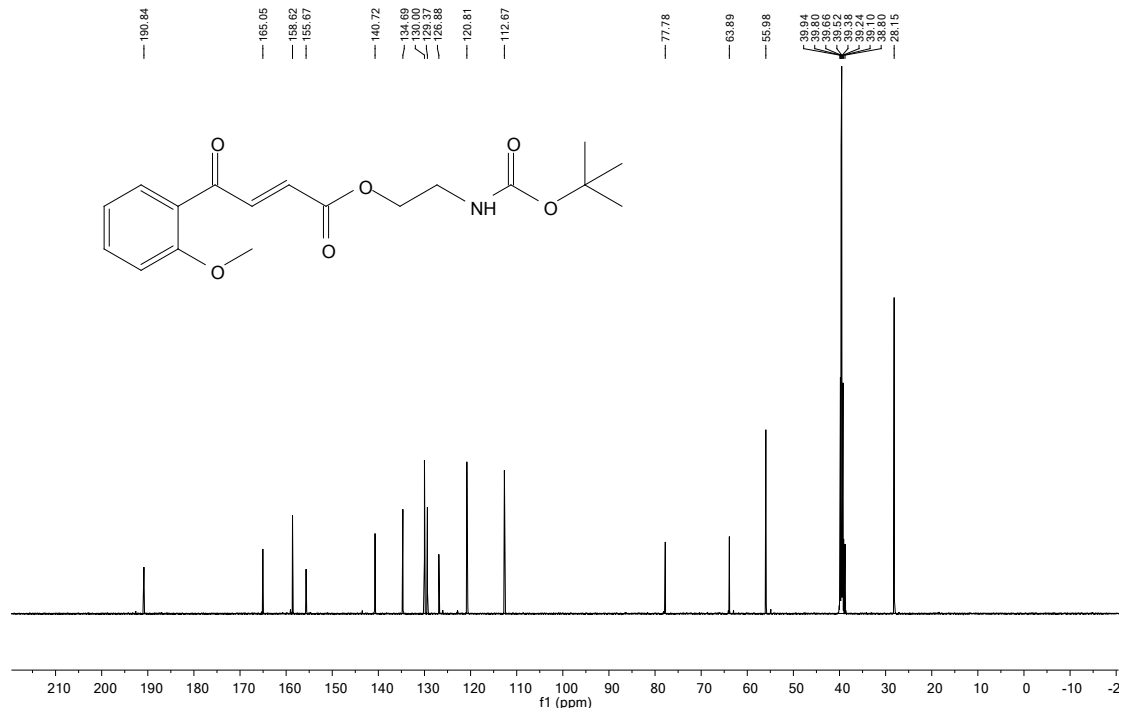
BRUKER AVANCE III 13C-NMR 11d IN DMSO-d6



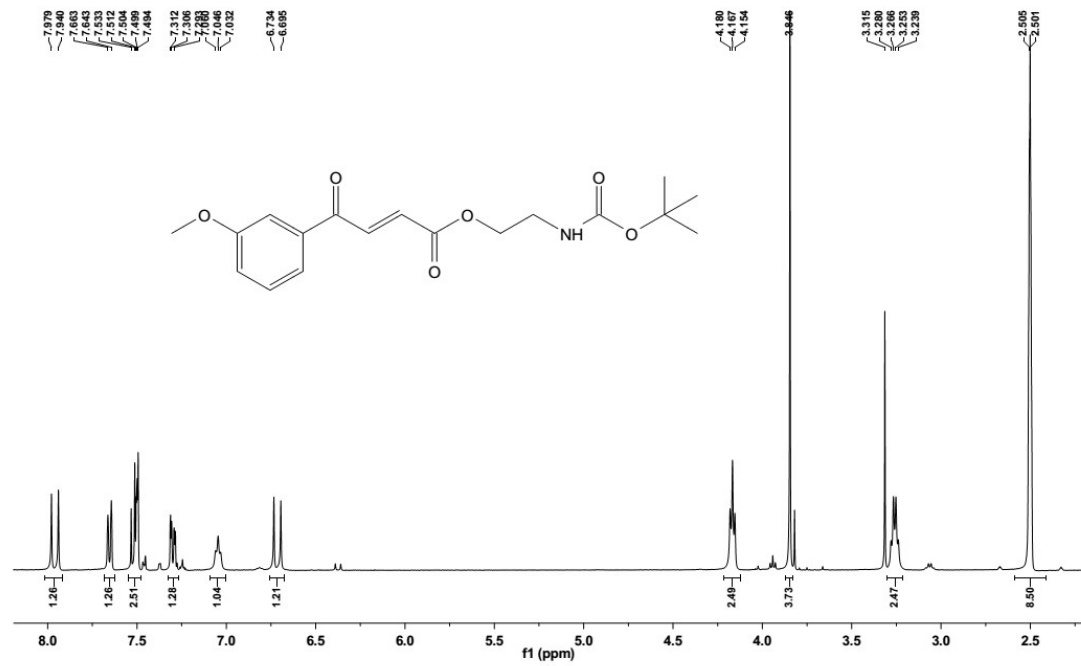
11e



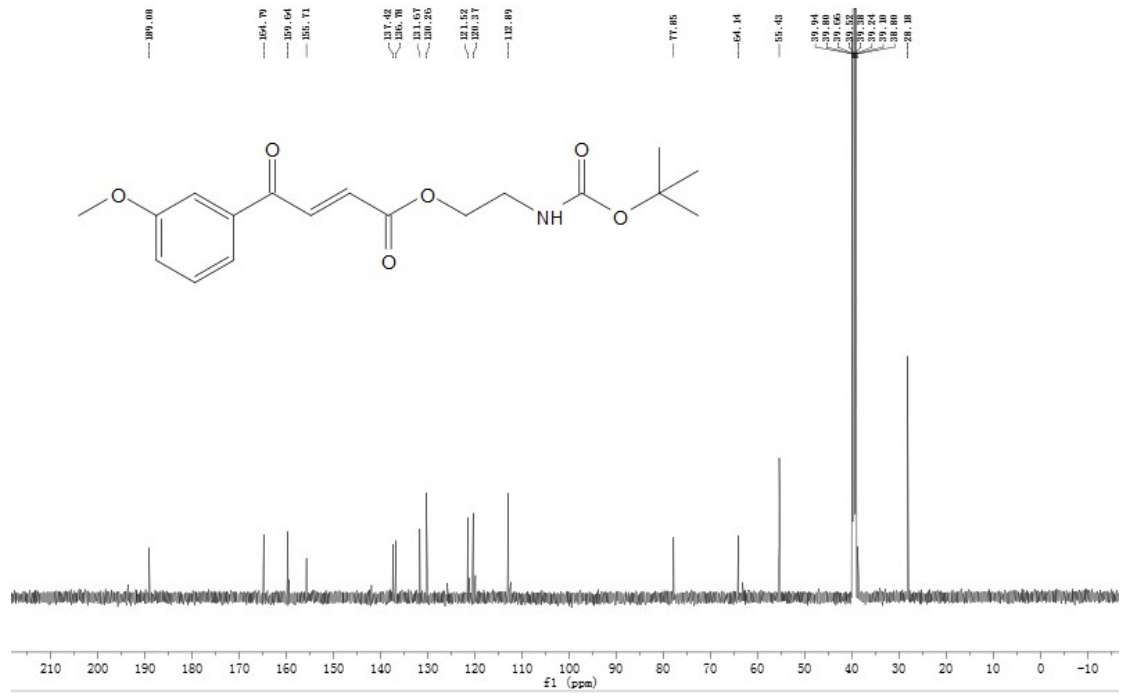
BRUKER AVANCE III 13C-NMR 11e IN DMSO-d6



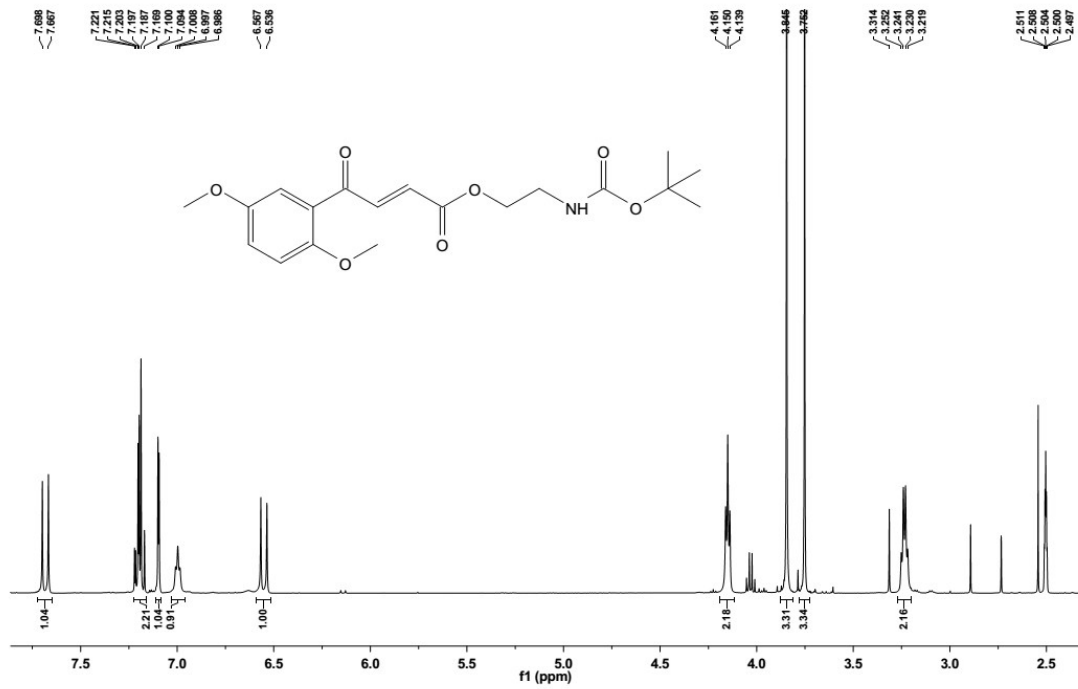
11f



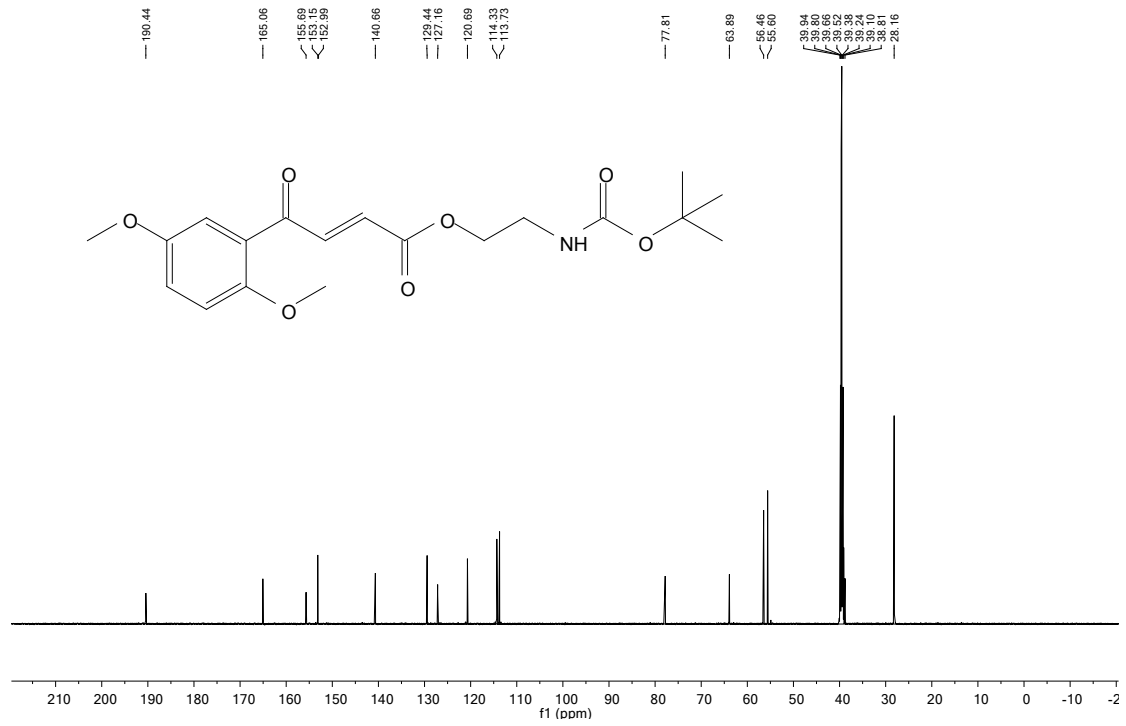
BRUKER AVANCE III 13C-NMR 11f IN DMSO-d6



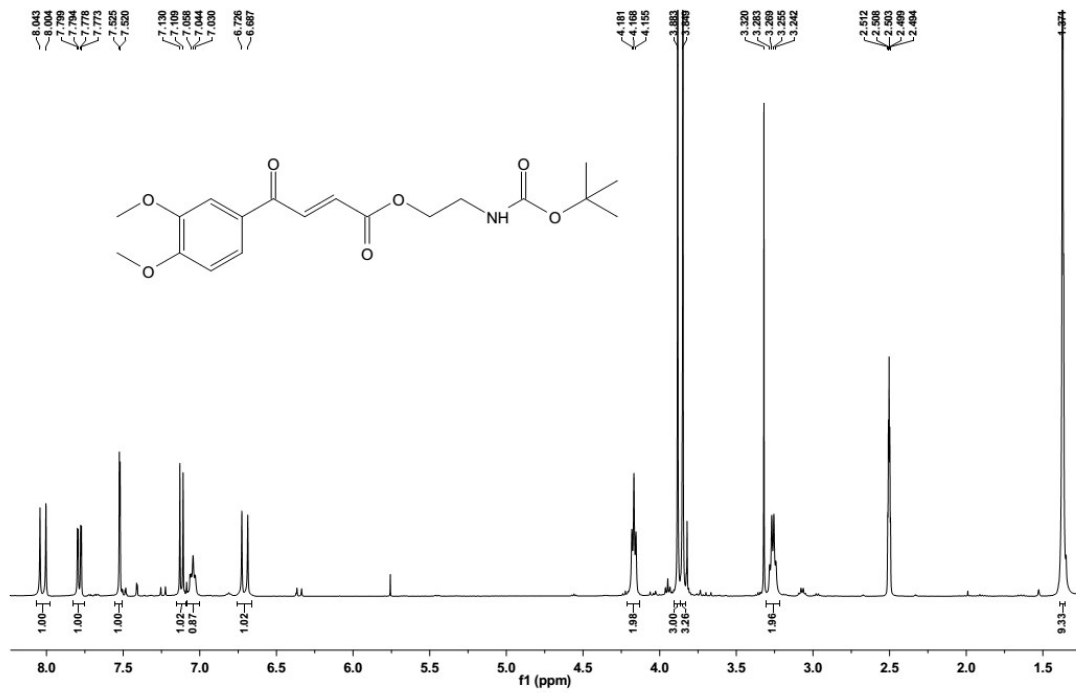
11g



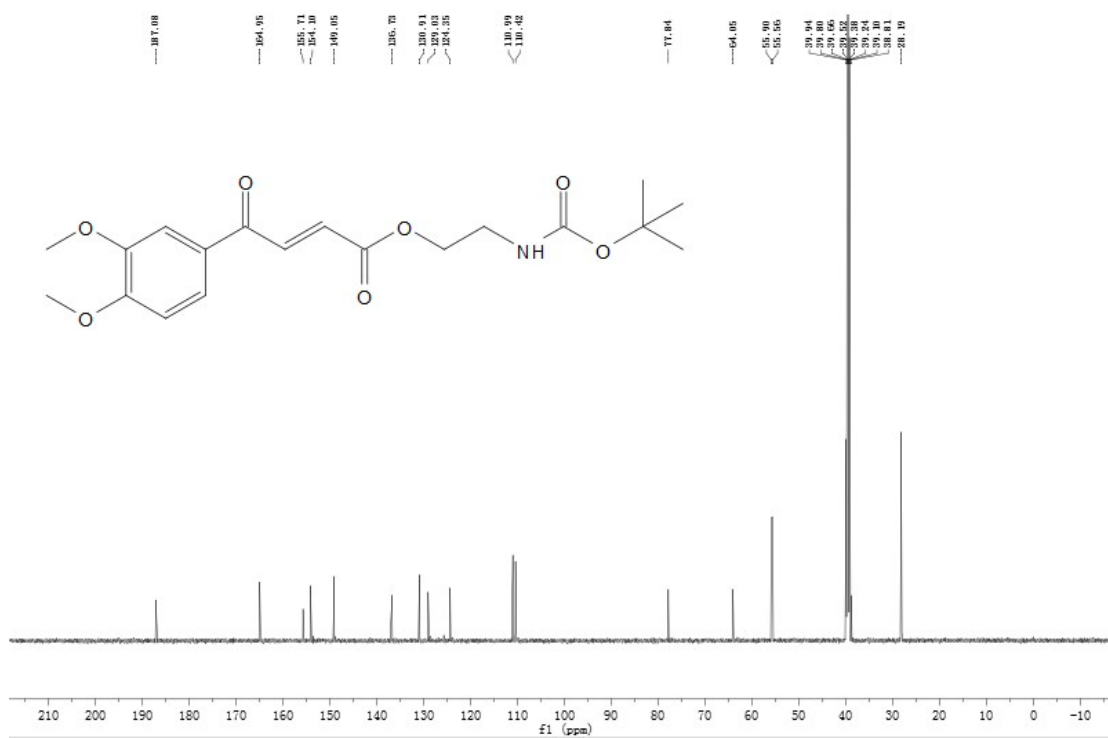
BRUKER AVANCE III 13C-NMR 11g IN DMSO-d6



11h

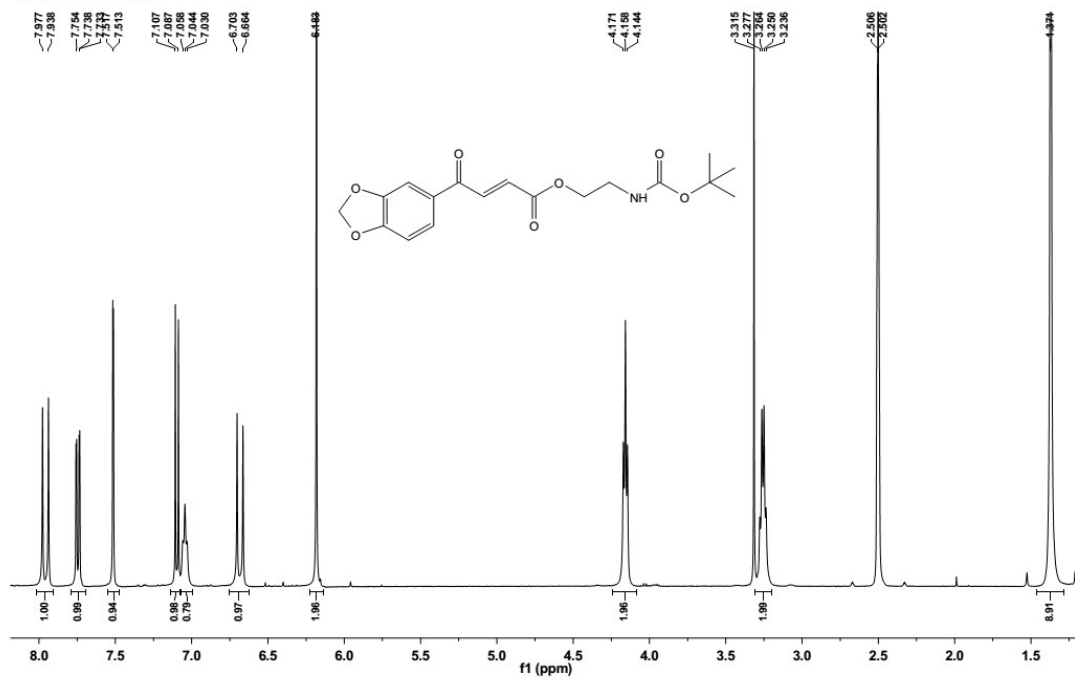


BRUKER AVANCE III 13C-NMR 11h IN DMSO-d6

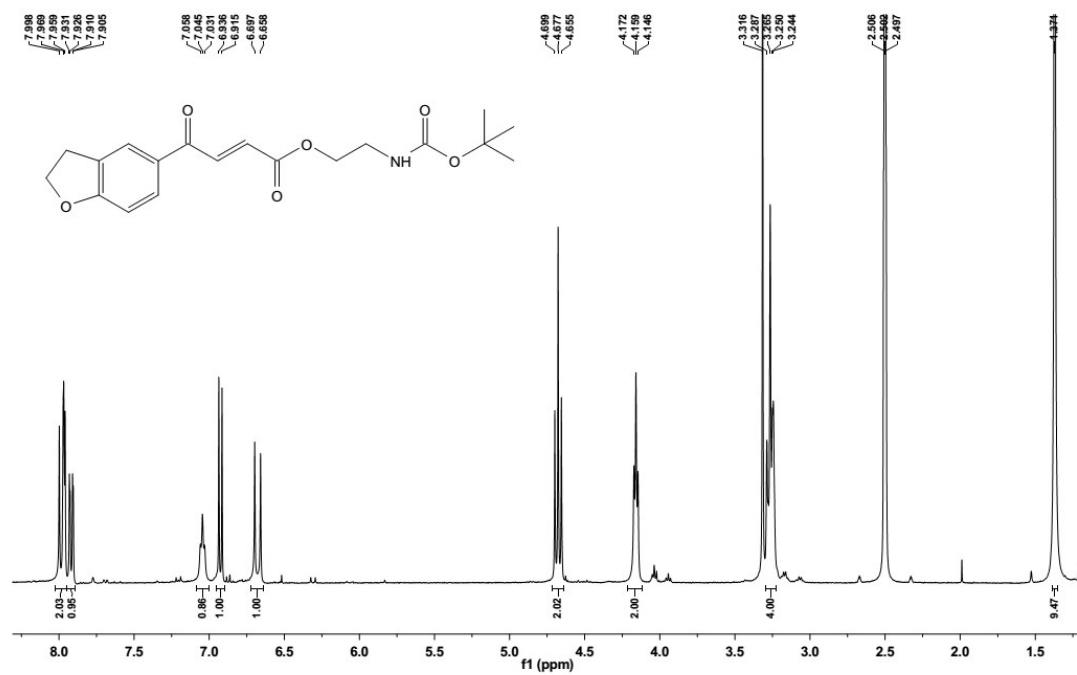


11i

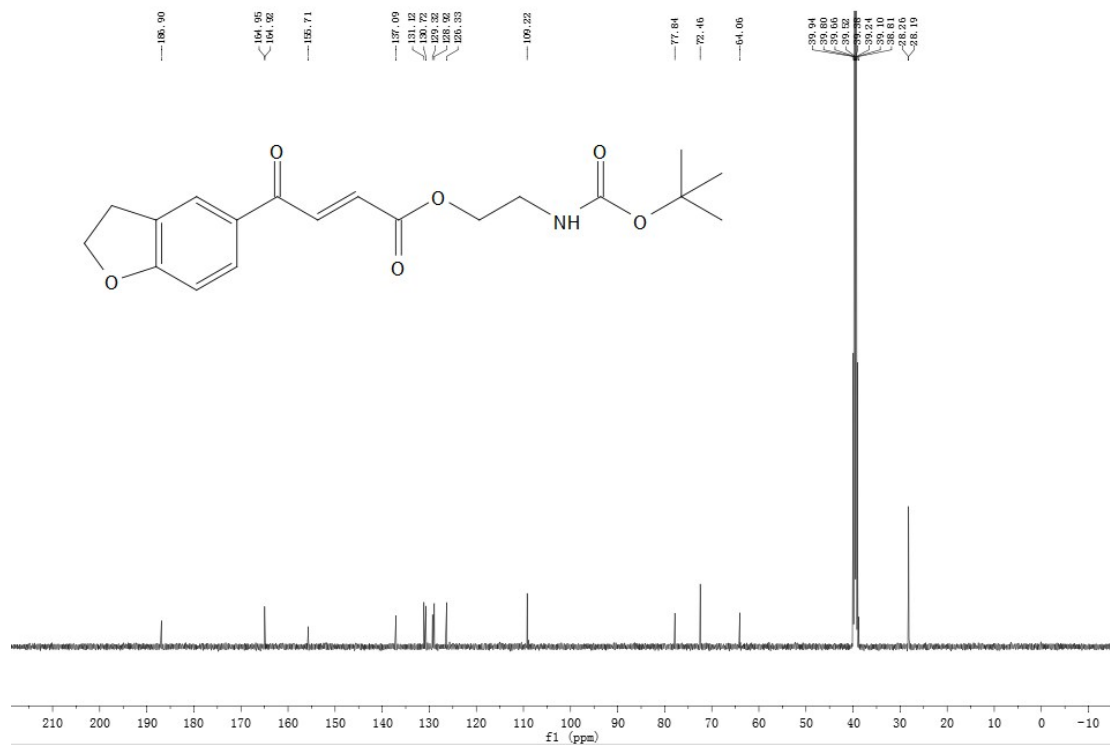
20140304 XCL13E35F
Bruker AVANCE III 400 20140304
PROTON2 DMSO



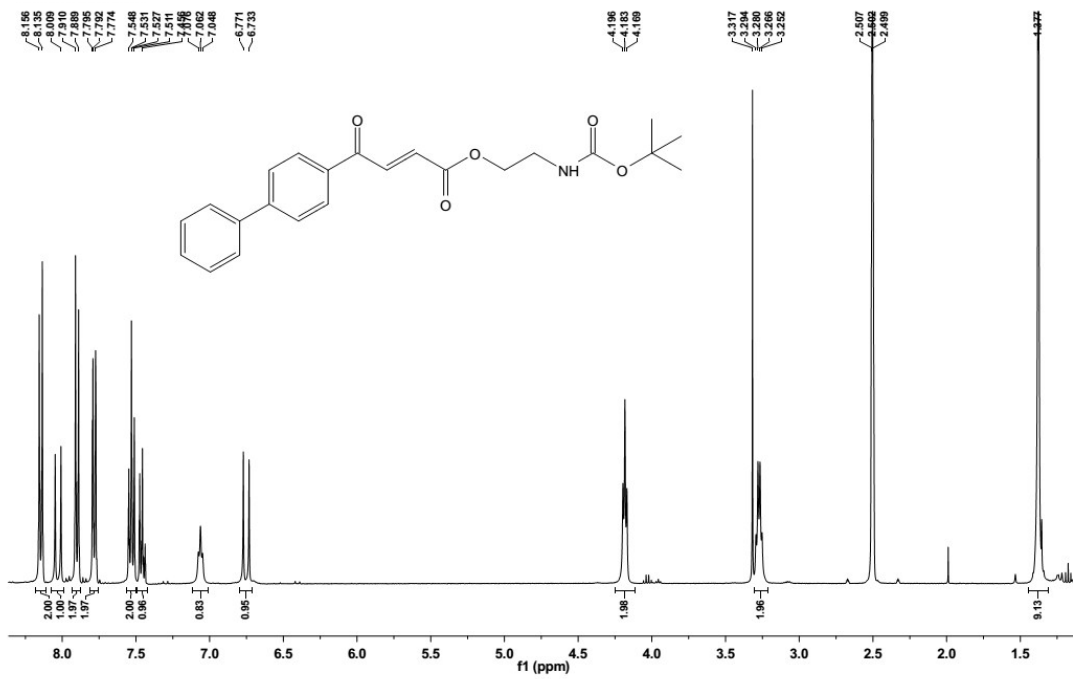
11j



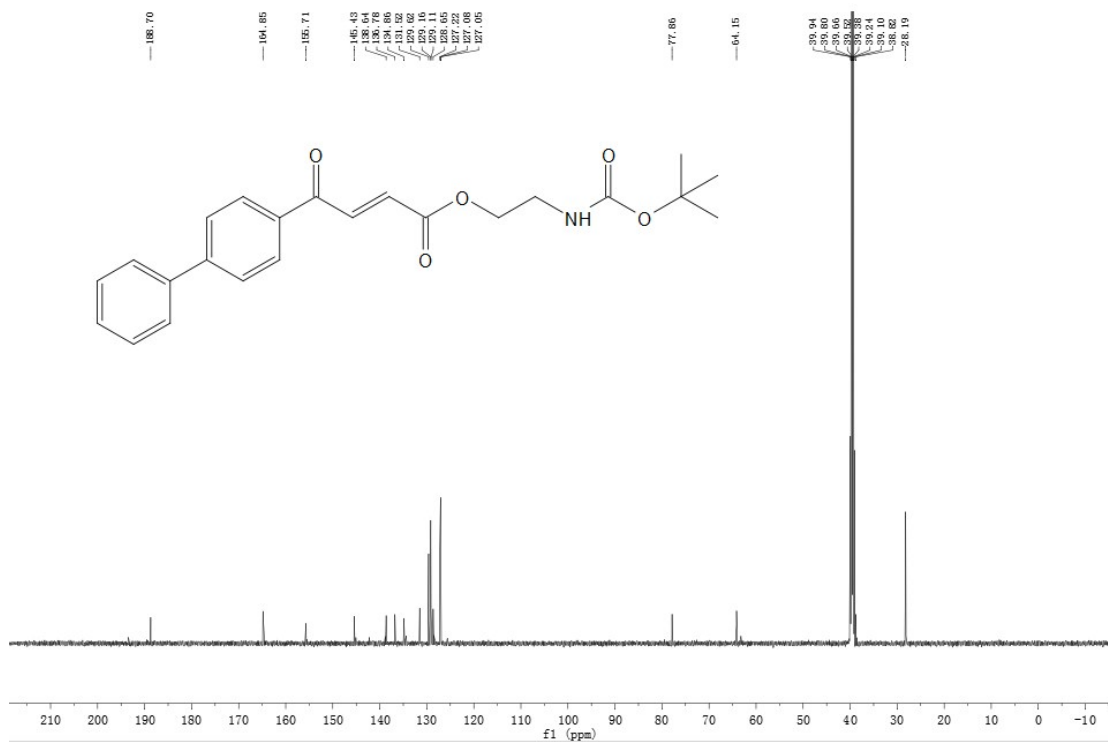
BRUKER AVANCE III 13C-NMR 11j IN DMSO-d6

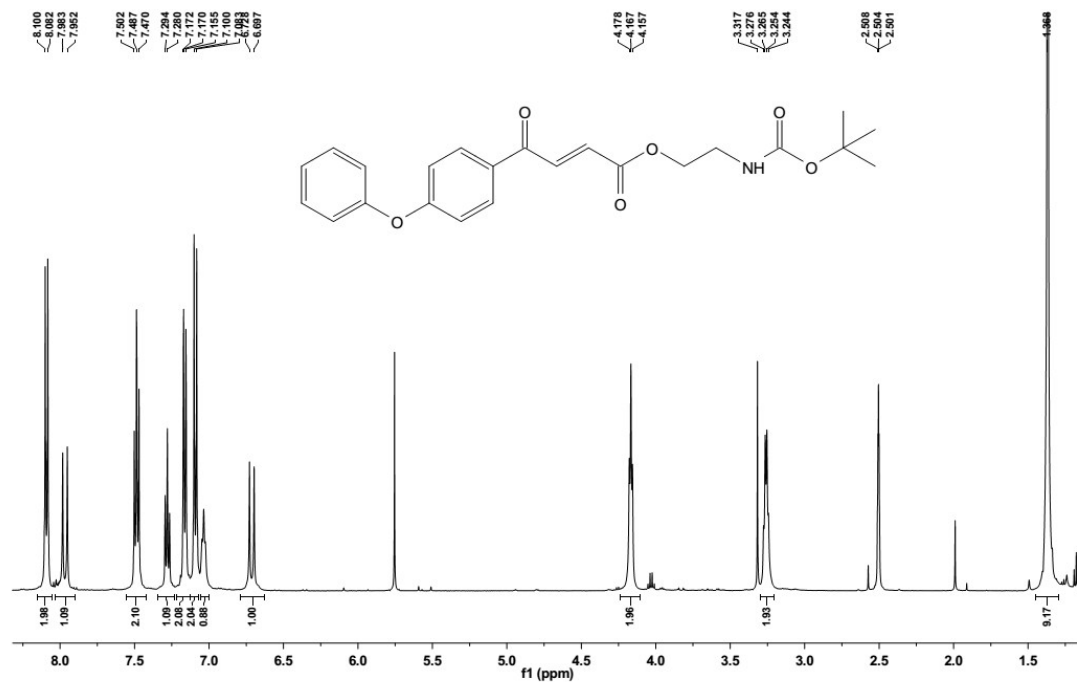


11k

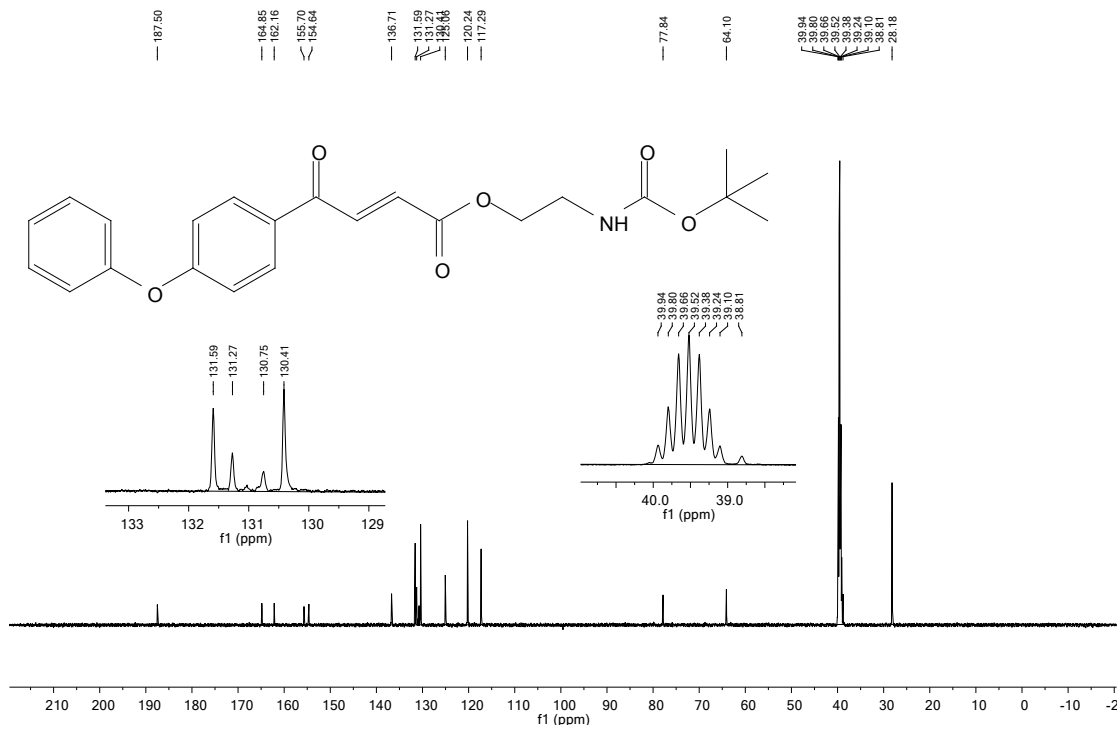


BRUKER AVANCE III 13C-NMR 11k IN DMSO-d6

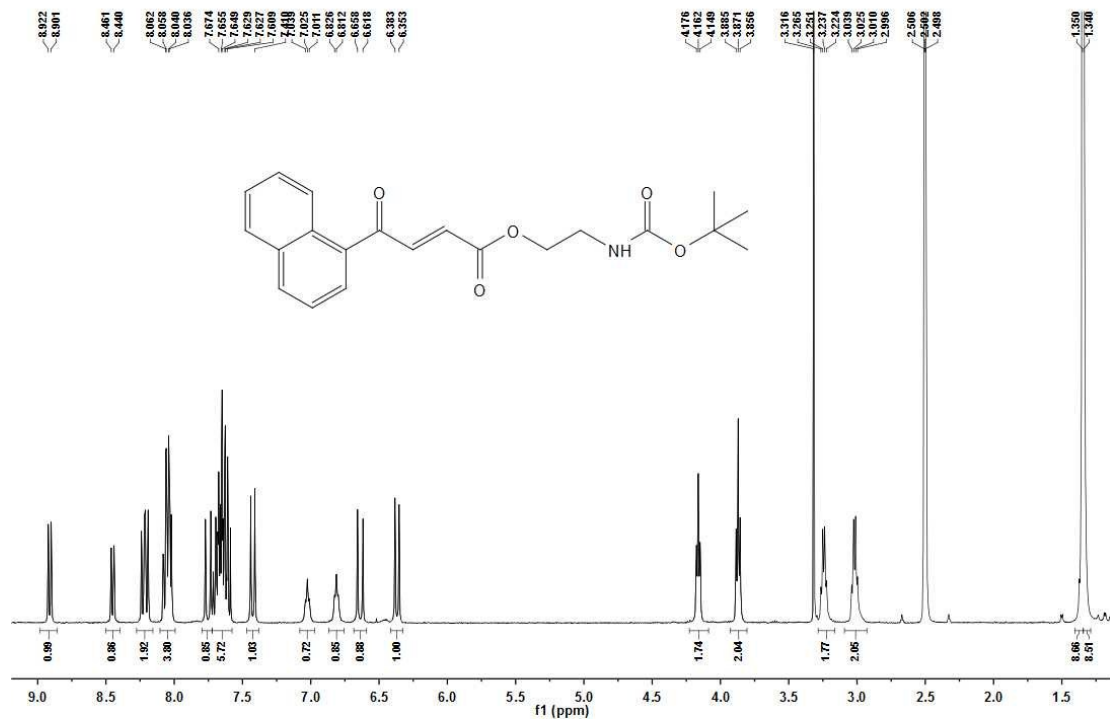




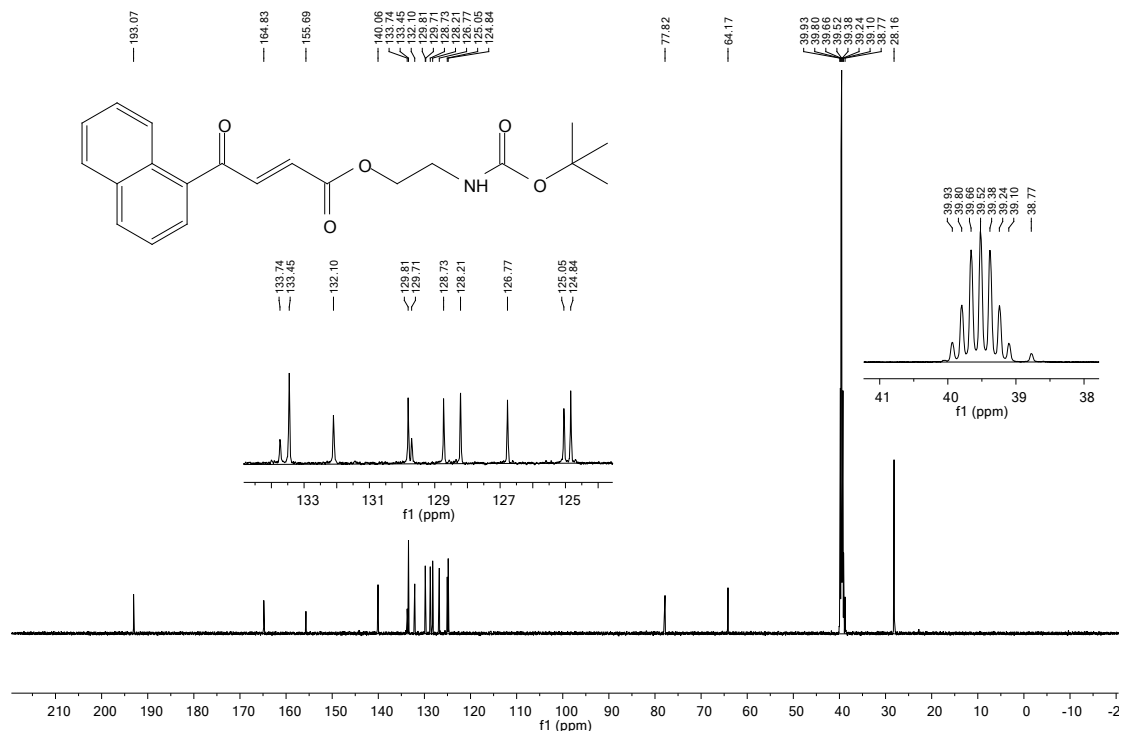
BRUKER AVANCE III 13C-NMR 111 IN DMSO-d6



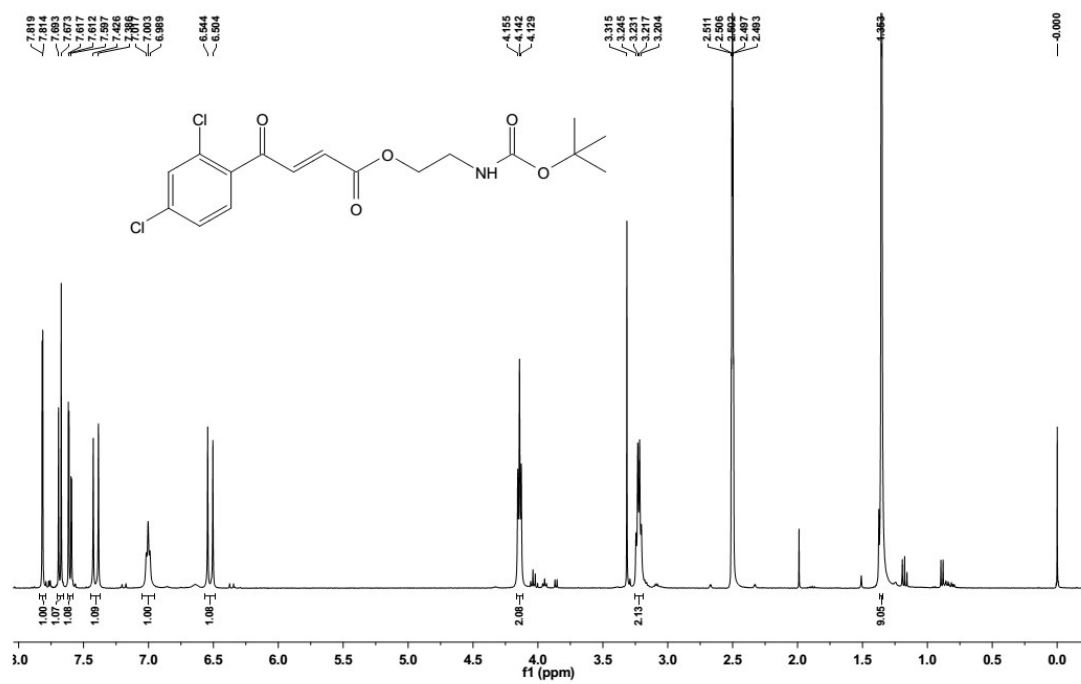
11m



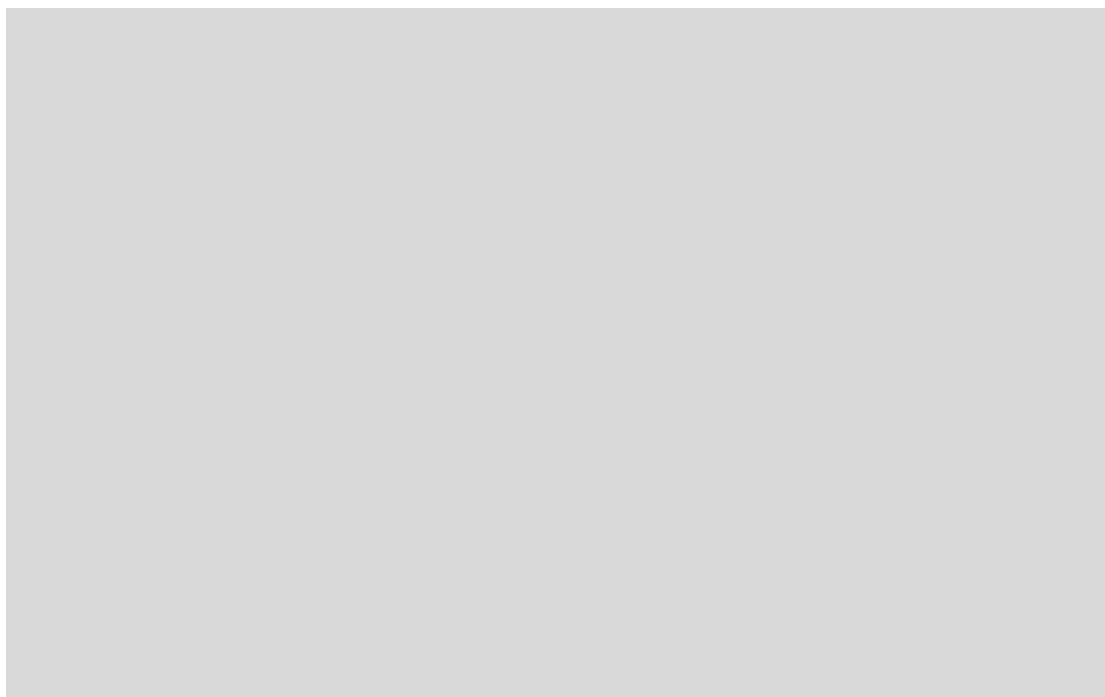
BRUKER AVANCE III 13C-NMR 11m IN DMSO-d6



11n



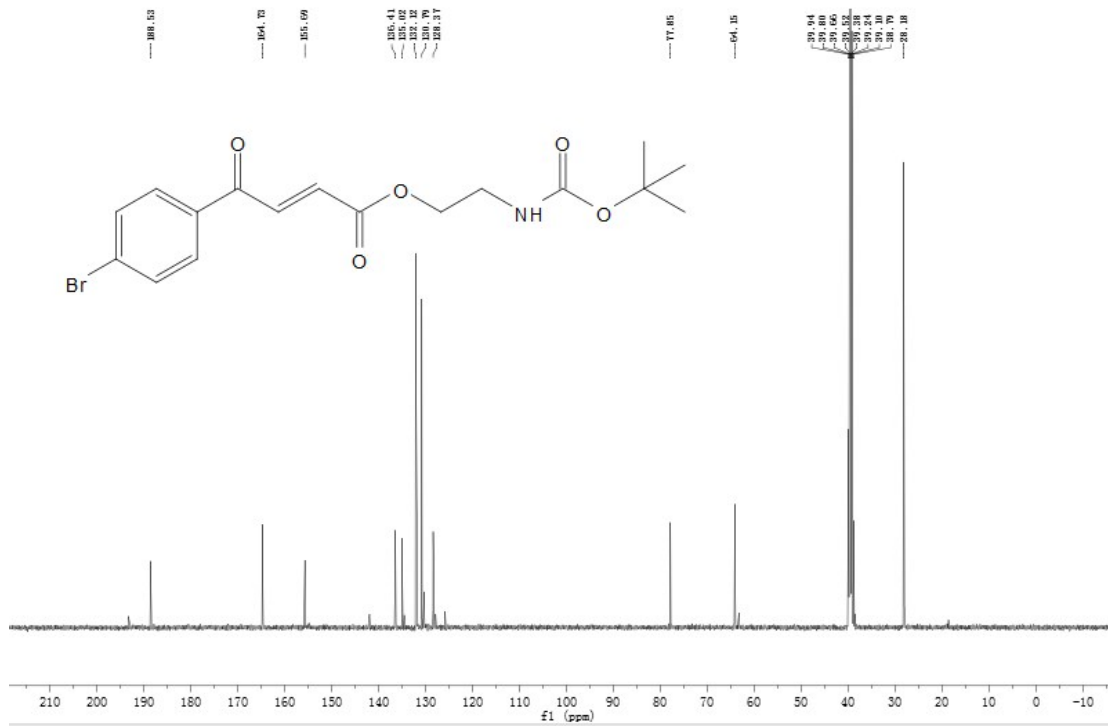
11o



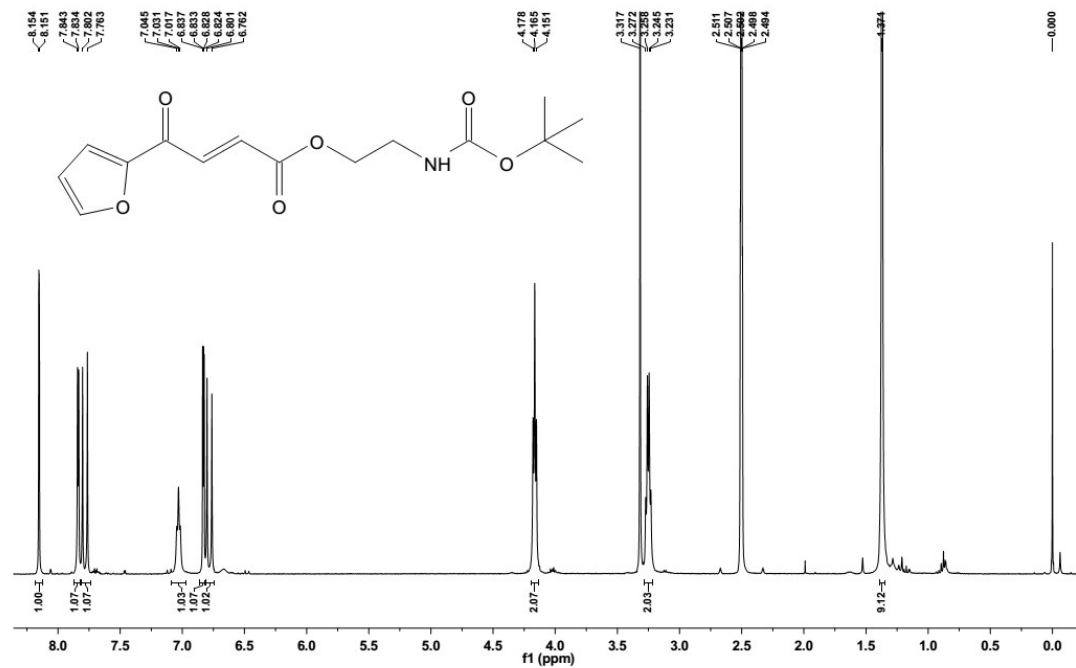
11p



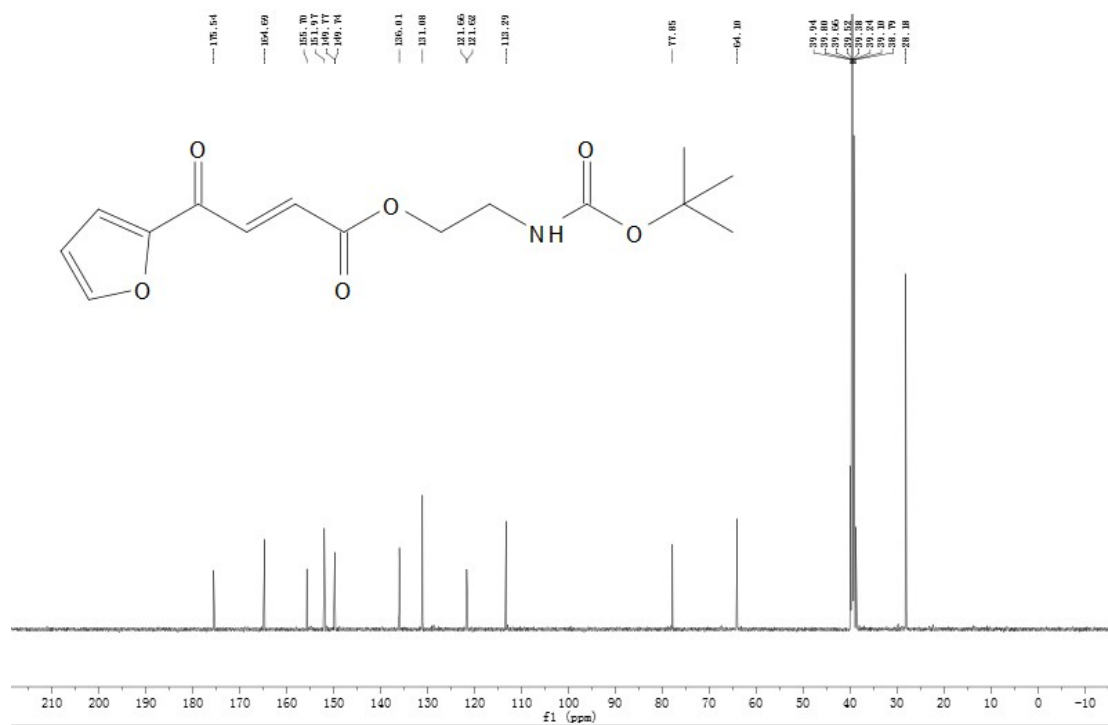
BRUKER AVANCE III 13C-NMR 11p IN DMSO-d6



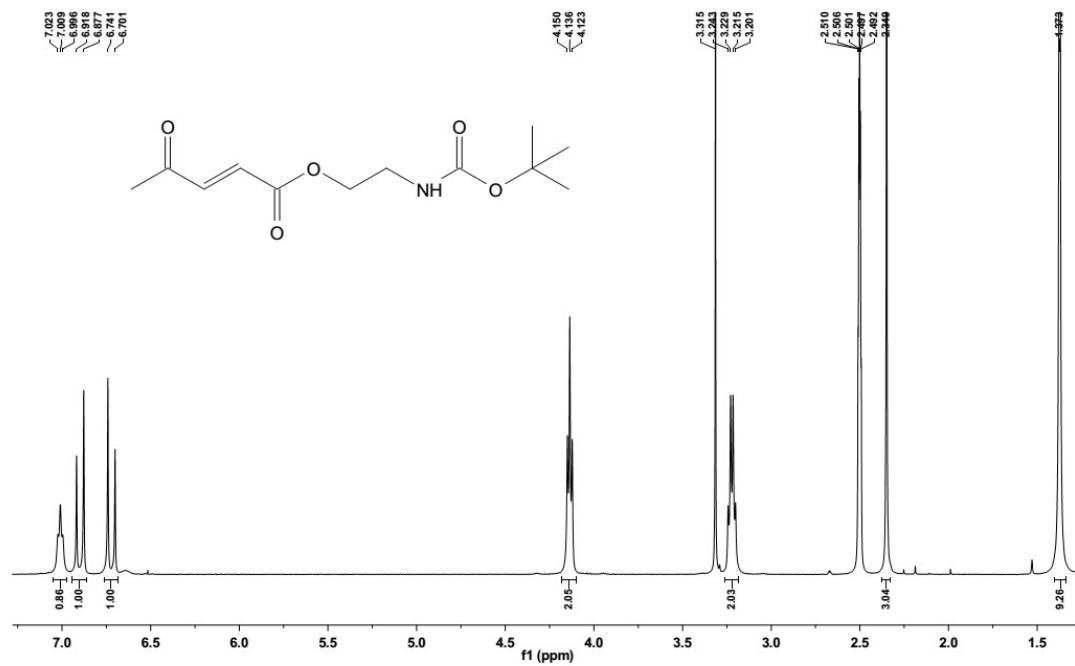
11q



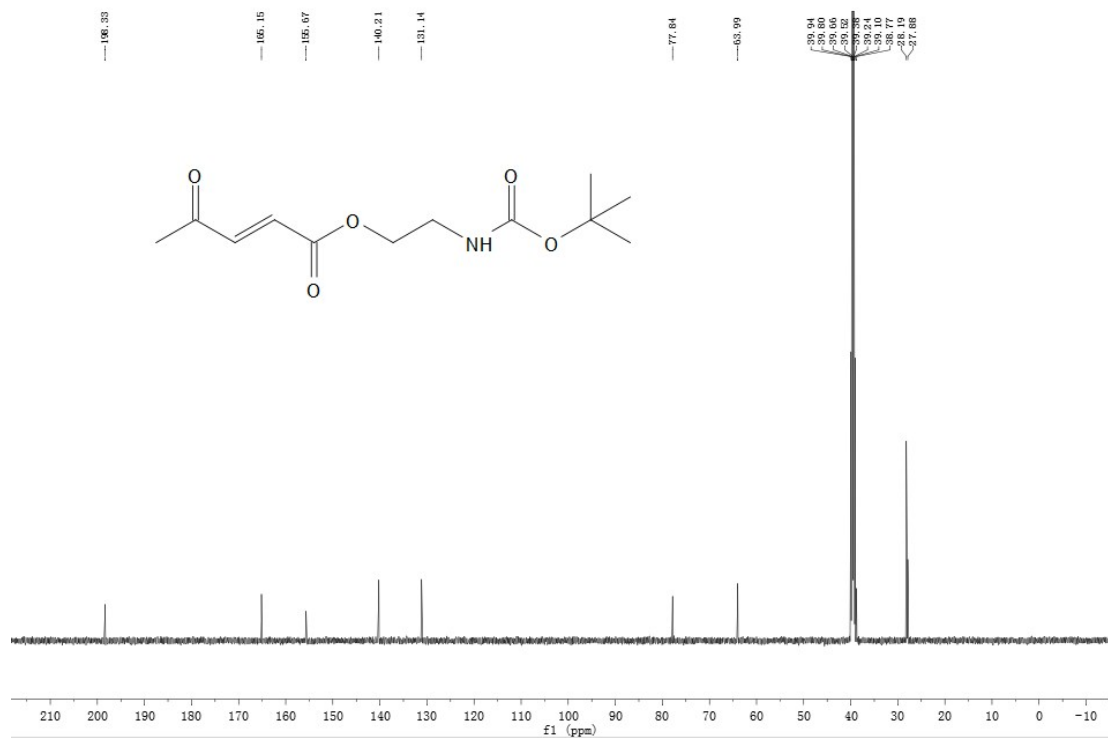
BRUKER AVANCE III 13C-NMR 11q IN DMSO-d6



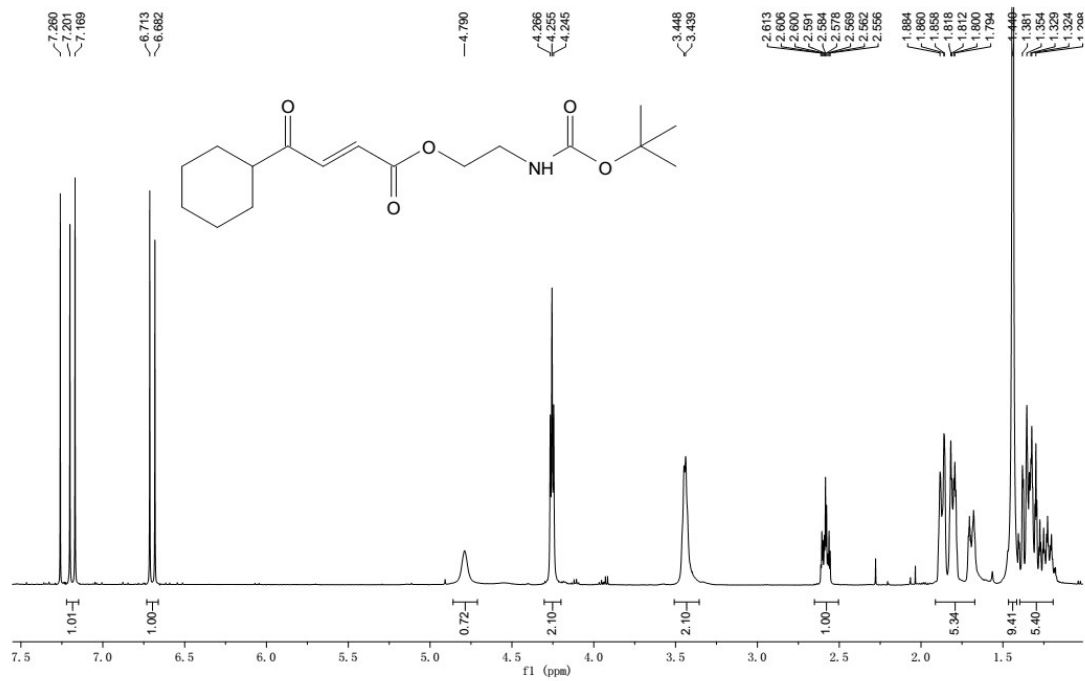
11r



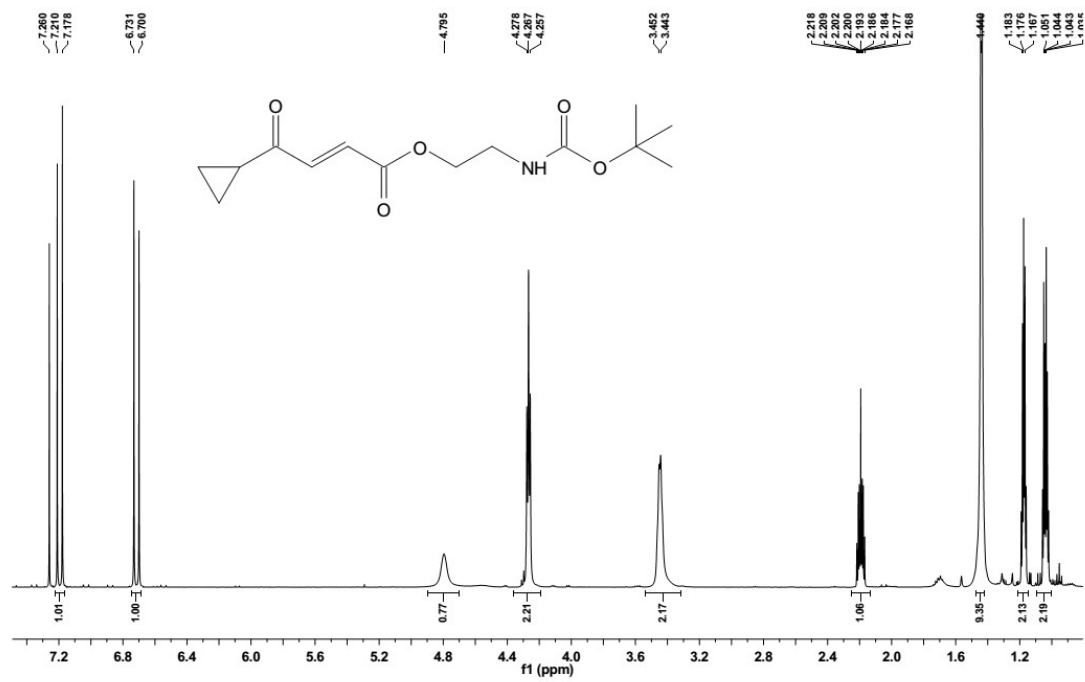
BRUKER AVANCE III 13C-NMR 11x IN DMSO-d6



11s



11t



BRUKER AVANCE III 13C-NMR 11t IN DMSO-d6

