

Supplementary Information

New acyl derivatives of 3-aminofurazanes and their antiplasmodial activities

Theresa Hermann ¹, Patrick Hochegger ^{1,*}, Johanna Dolensky ¹, Werner Seebacher ¹, Robert Saf ², Marcel Kaiser ³, Pascal Mäser ³ and Robert Weis ¹

¹ Institute of Pharmaceutical Sciences, Pharmaceutical Chemistry, University of Graz, Schubertstraße 1, A-8010 Graz, Austria; theresa.hermann@uni-graz.at; patrick.hochegger@uni-graz.at; we.seebacher@uni-graz.at; robert.weis@uni-graz.at

² Institute for Chemistry and Technology of Materials (ICTM), Graz University of Technology, Stremayrgasse 9, A-8010 Graz, Austria; robert.saf@tugraz.at

³ Swiss Tropical and Public Health Institute, Socinstraße 57, CH-4002 Basel, Switzerland; marcel.kaiser@swisstph.ch; pascal.maeser@swisstph.ch

* Correspondence: patrick.hochegger@uni-graz.at; Tel.: +43-316-380-5379; fax: +43-316-380-9846

NMR spectra data of compounds 6, 2, 8, 9, 20, 1, 10-17, 26-29, 31, 32, 34, 35 and 39

Figure S1. ^1H NMR at 400 MHz and ^{13}C NMR at 100 MHz spectra for compound 6

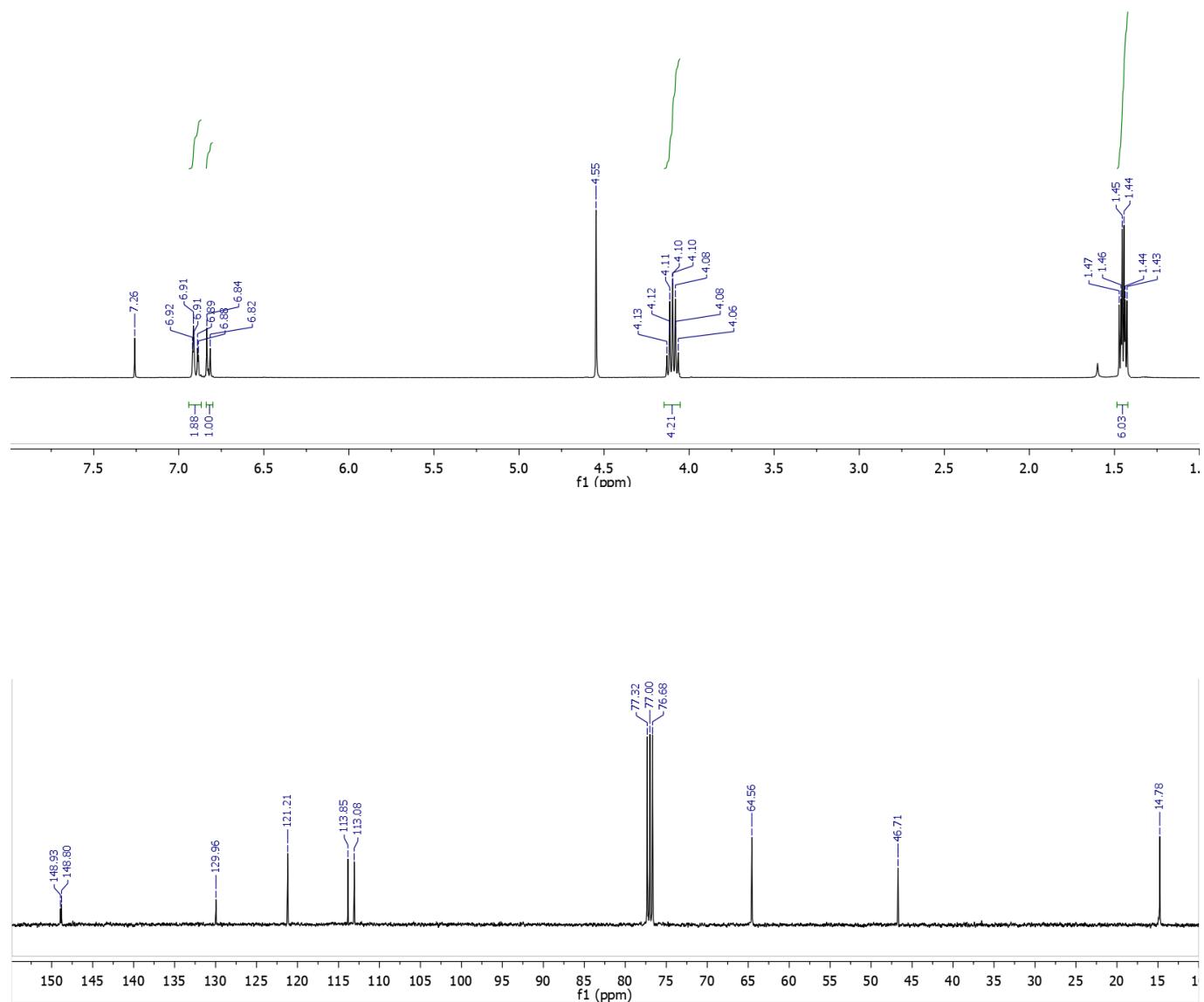


Figure S2. ^1H NMR at 400 MHz and ^{13}C NMR at 100 MHz spectra for compound 2

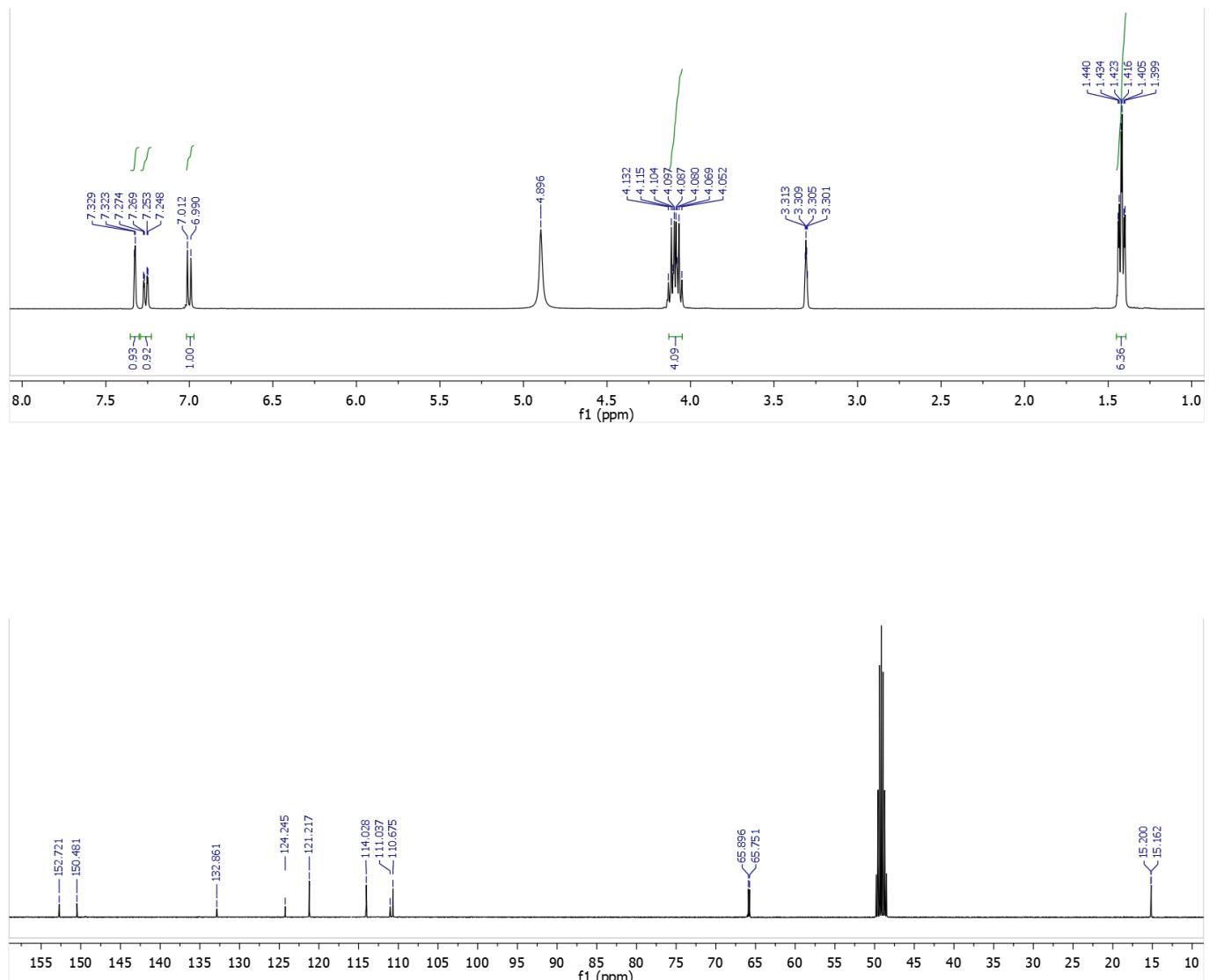


Figure S3. ^1H NMR at 400 MHz and ^{13}C NMR at 100 MHz spectra for compound 8

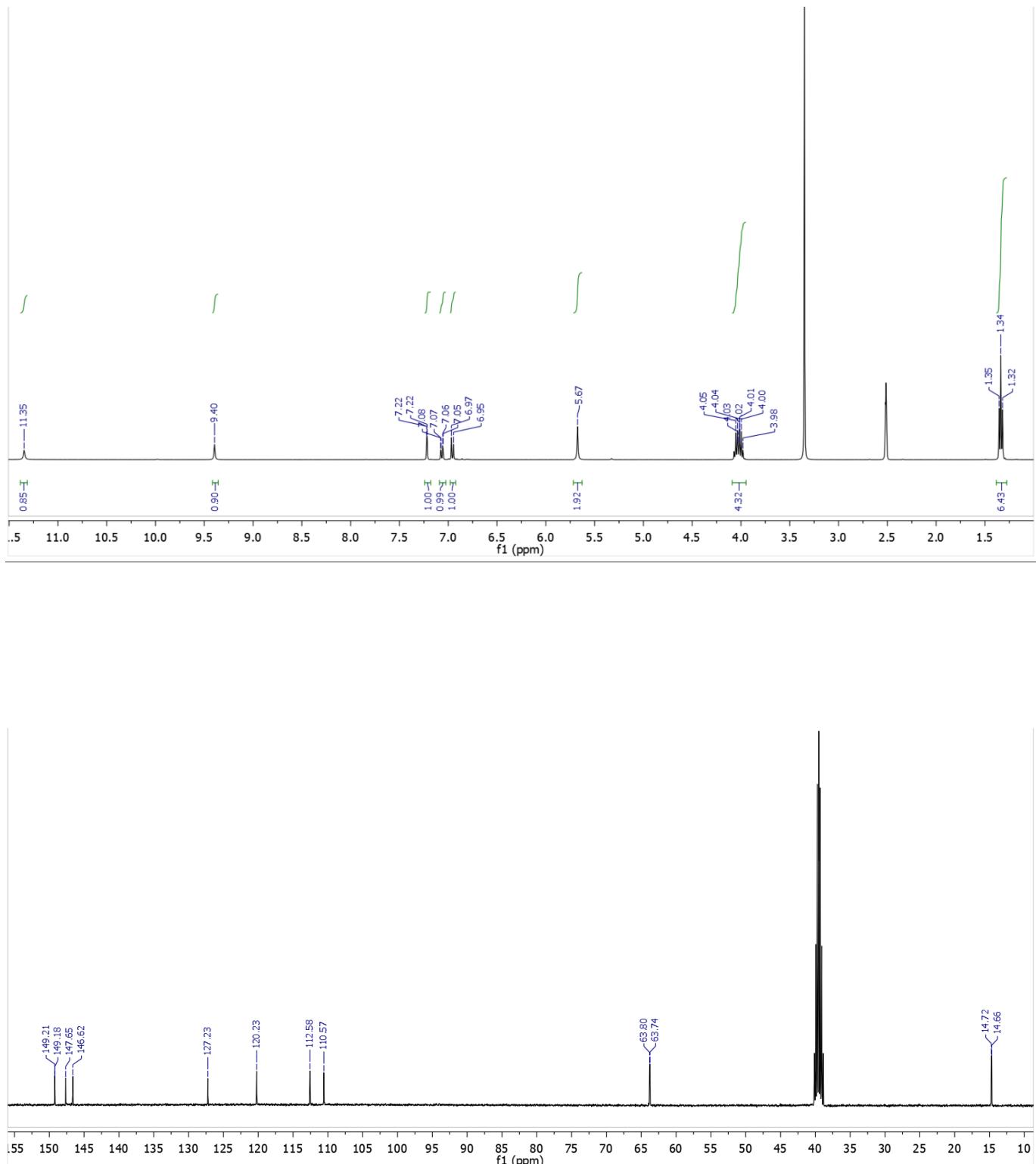


Figure S4. ^1H NMR at 400 MHz and ^{13}C NMR at 100 MHz spectra for compound 9

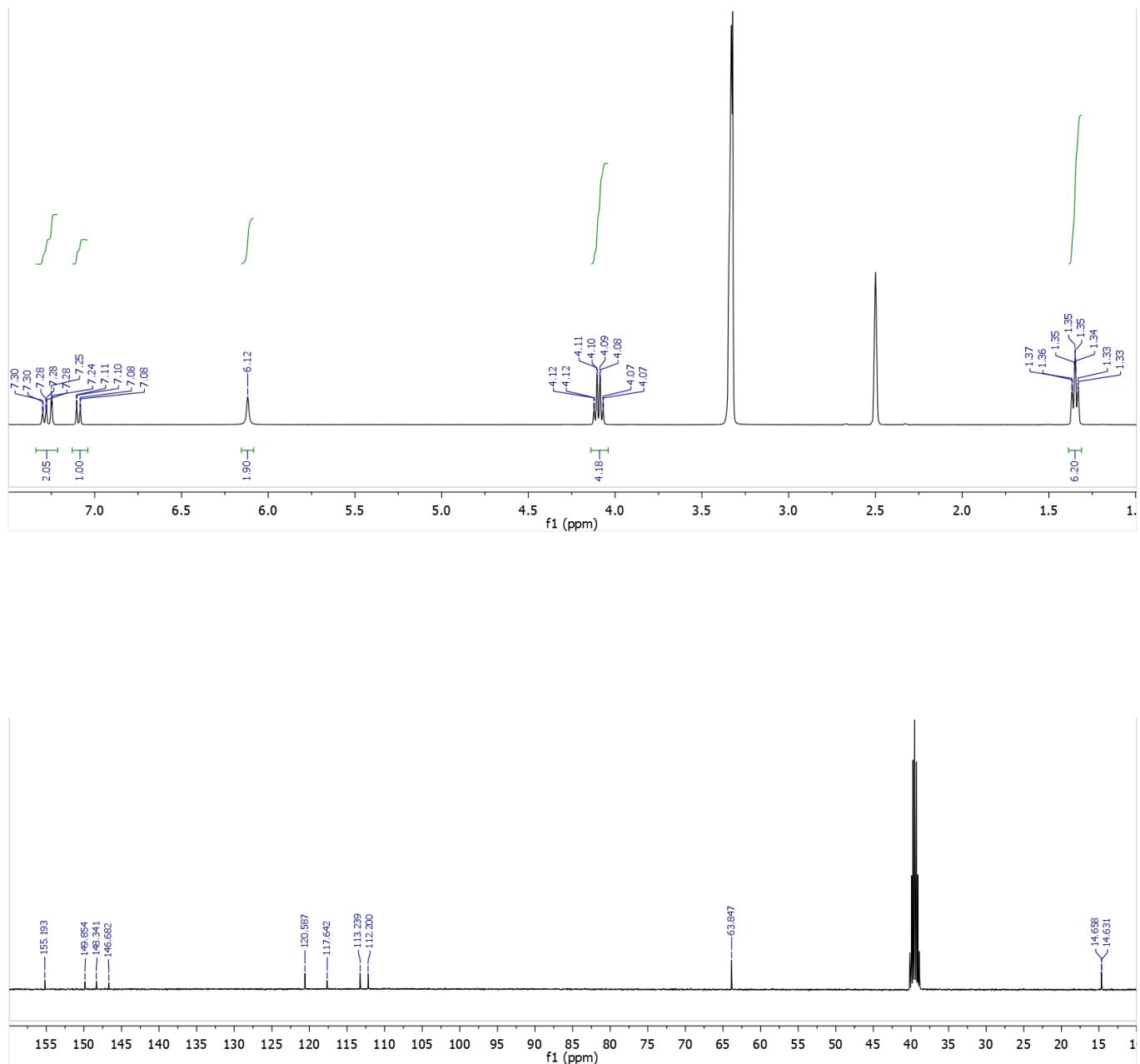


Figure S5. ^1H NMR at 400 MHz and ^{13}C NMR at 100 MHz spectra for compound **20**

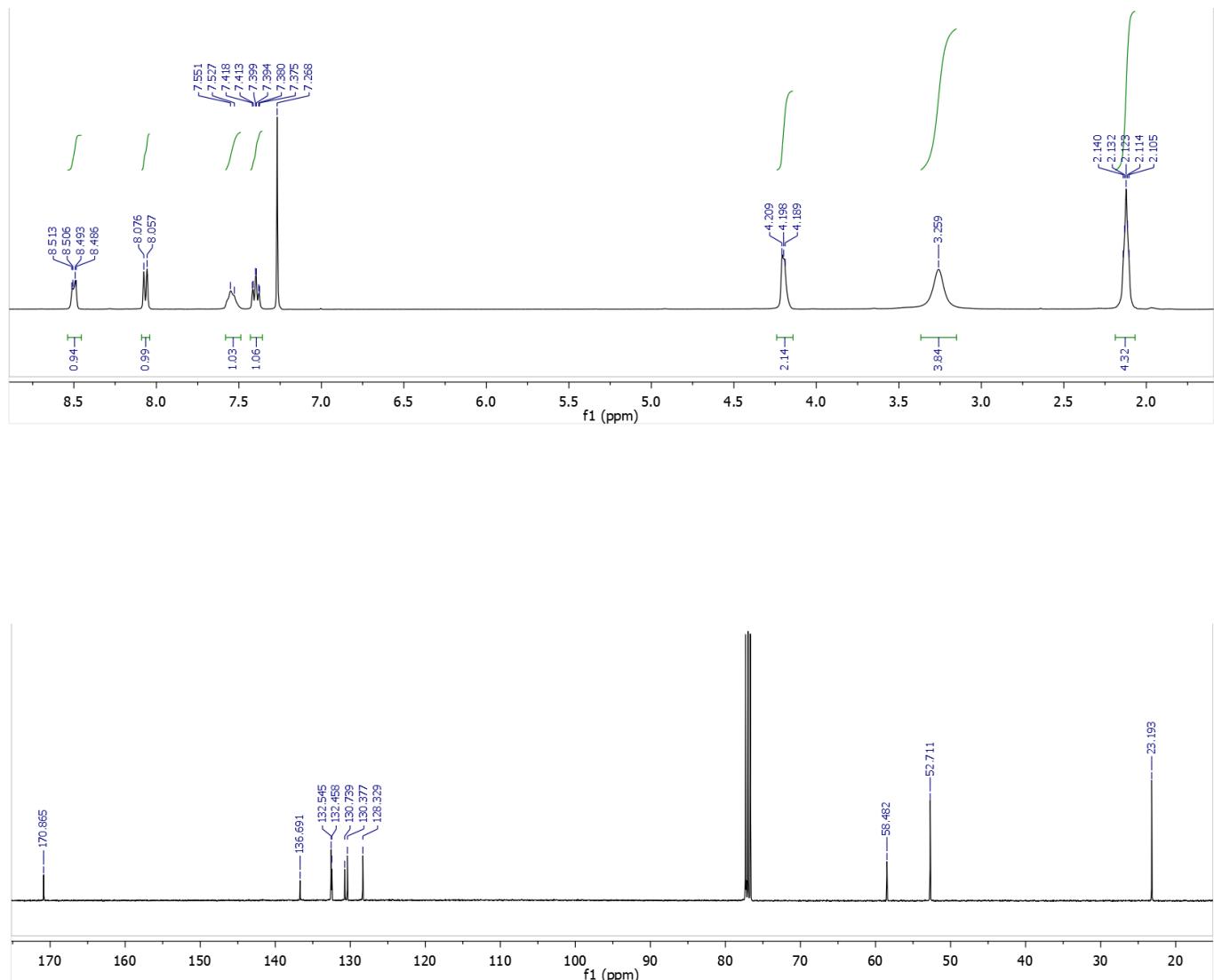


Figure S6. ^1H NMR at 400 MHz and ^{13}C NMR at 100 MHz spectra for compound **1**

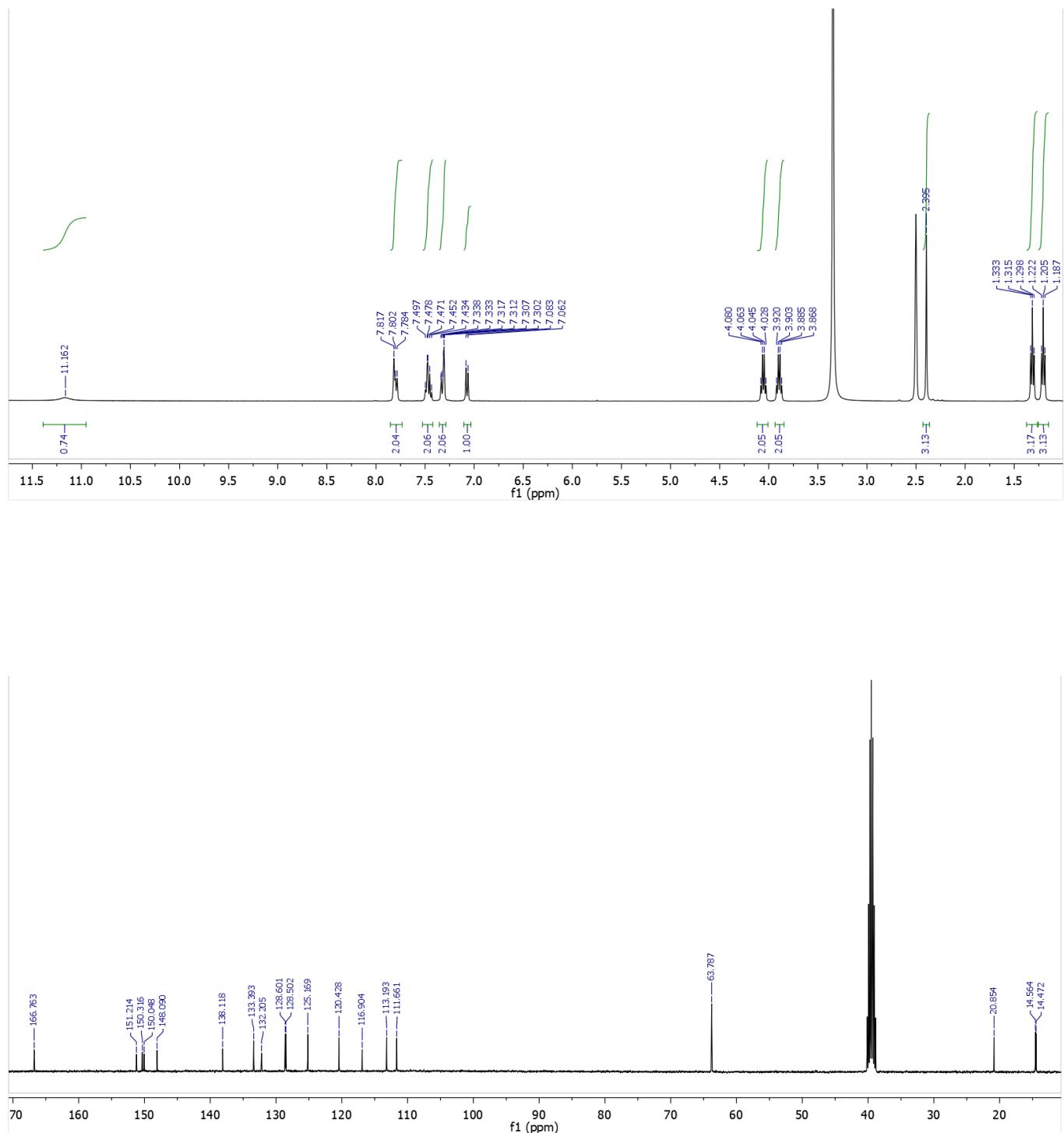


Figure S7. ^1H NMR at 400 MHz and ^{13}C NMR at 100 MHz spectra for compound **10**

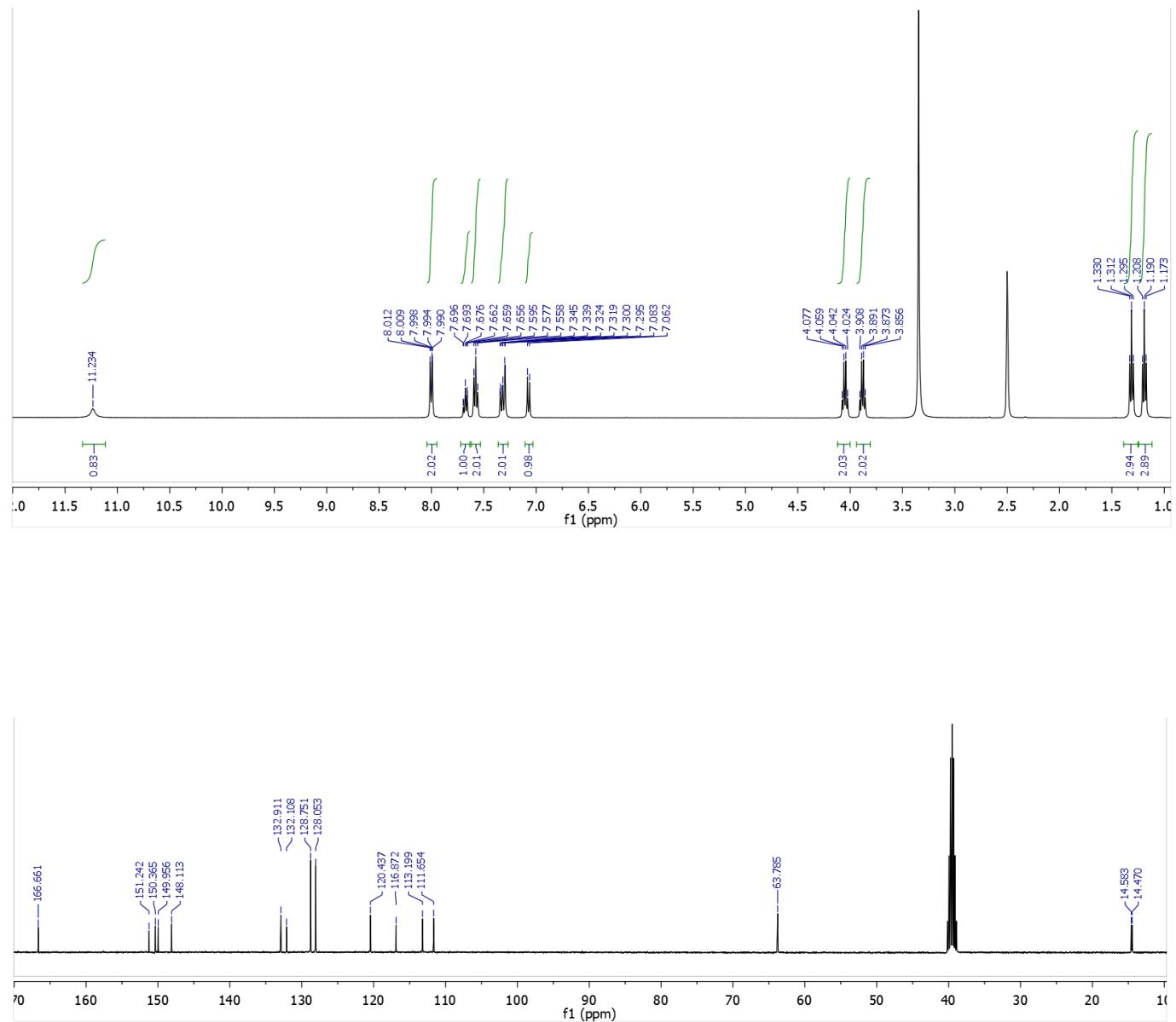


Figure S8. ^1H NMR at 400 MHz and ^{13}C NMR at 100 MHz spectra for compound 11

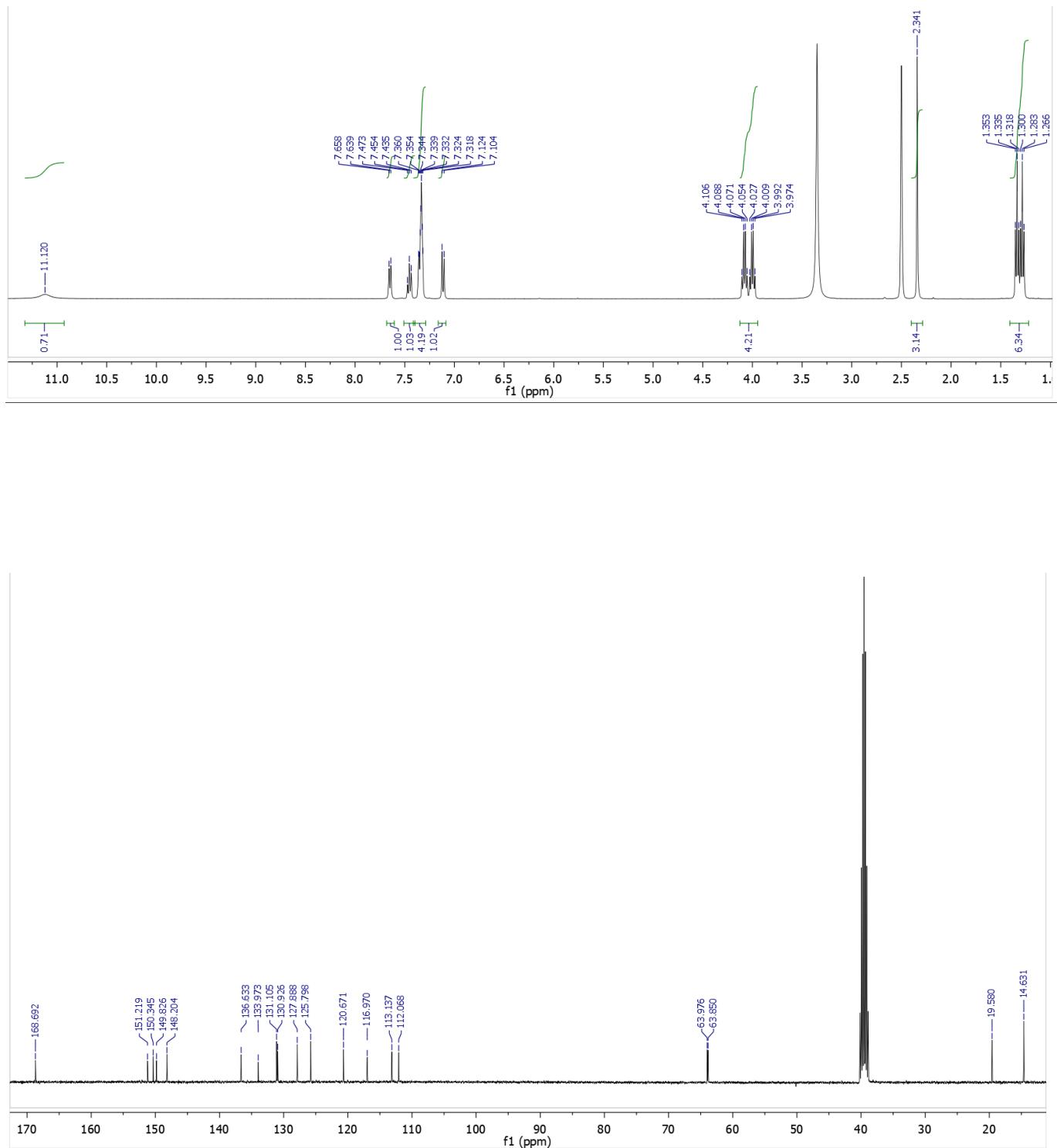


Figure S9. ^1H NMR at 400 MHz and ^{13}C NMR at 100 MHz spectra for compound 12

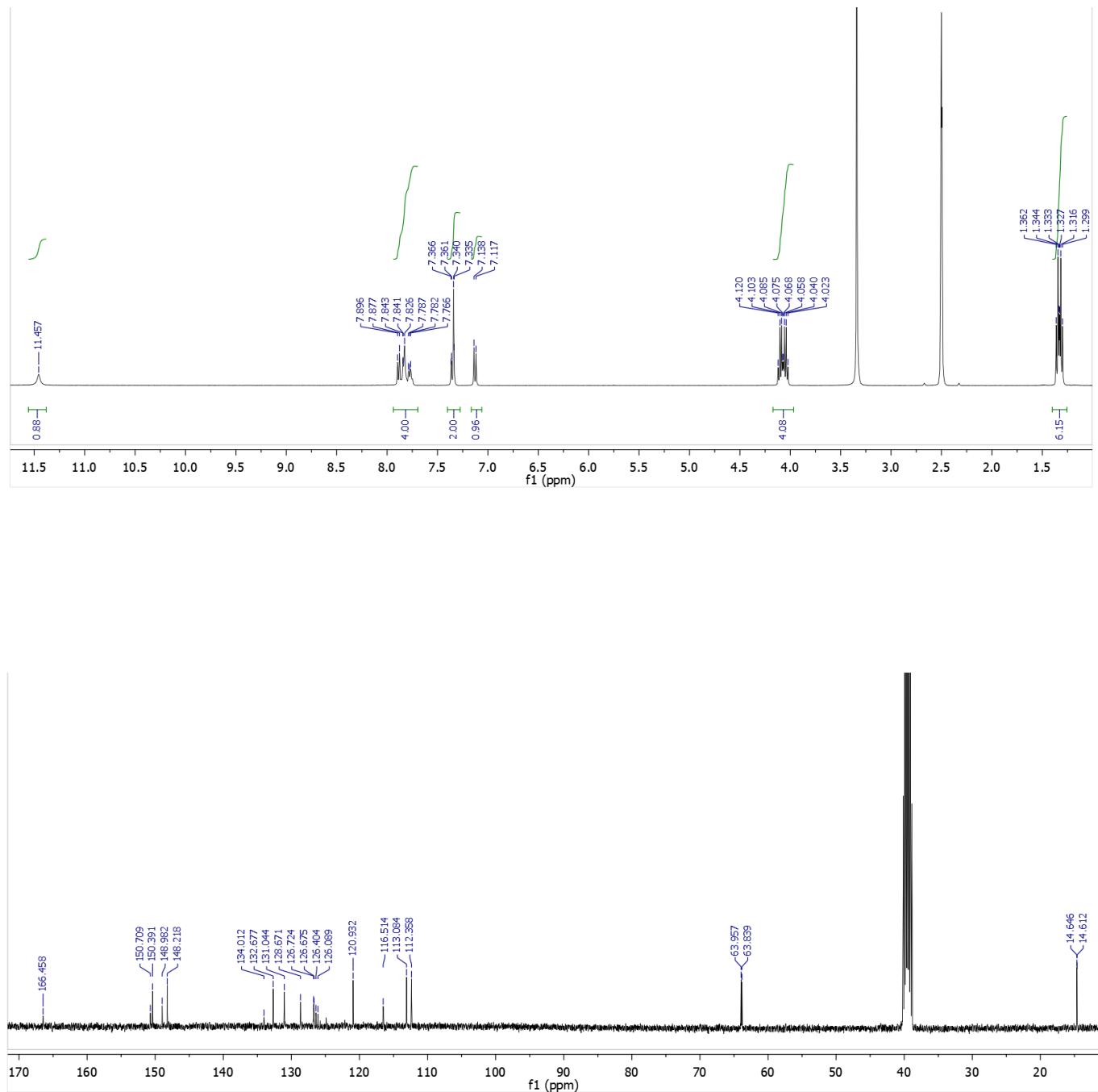


Figure S10. ^1H NMR at 400 MHz and ^{13}C NMR at 100 MHz spectra for compound **13**

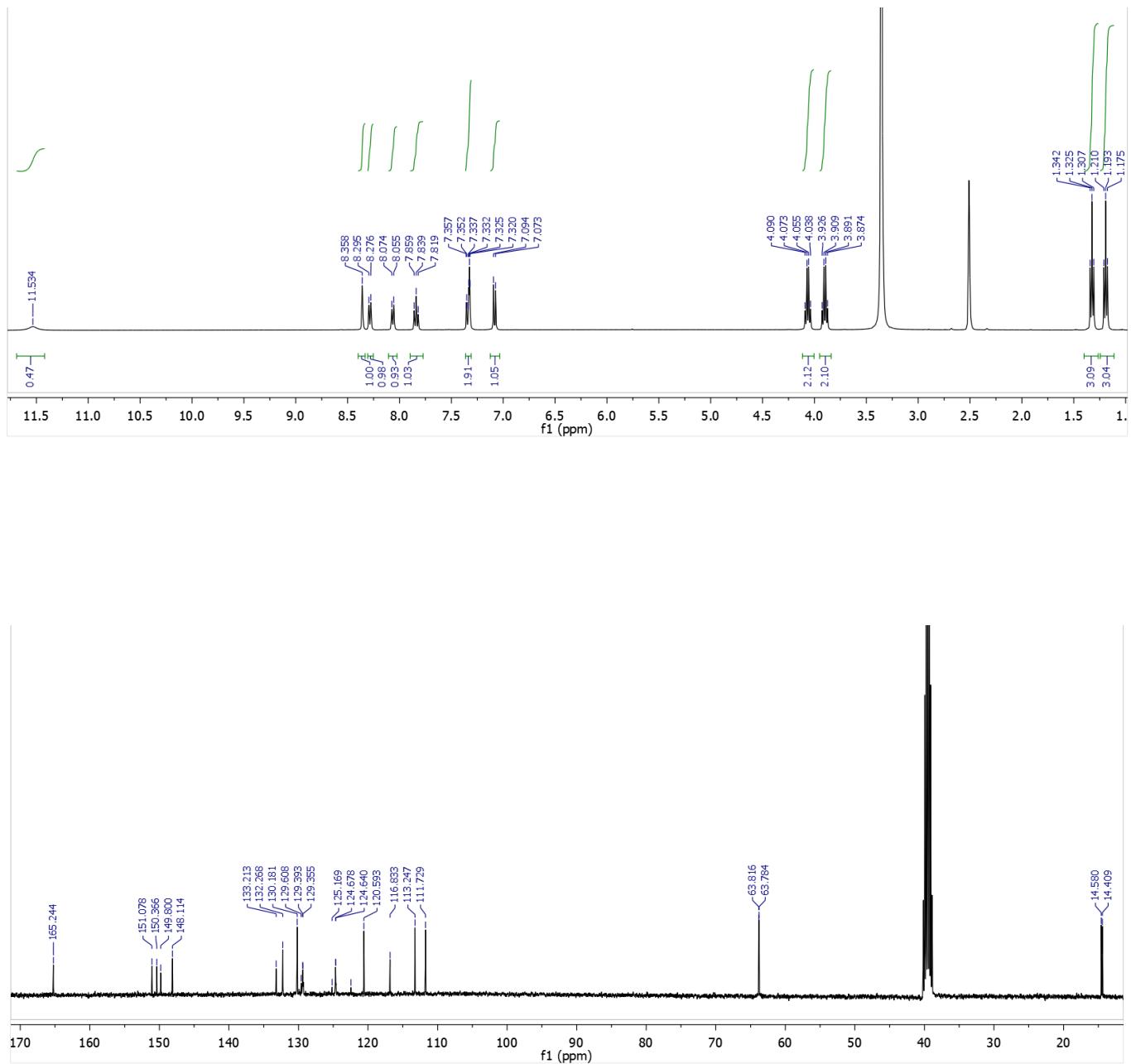


Figure S11. ^1H NMR at 400 MHz and ^{13}C NMR at 100 MHz spectra for compound **14**

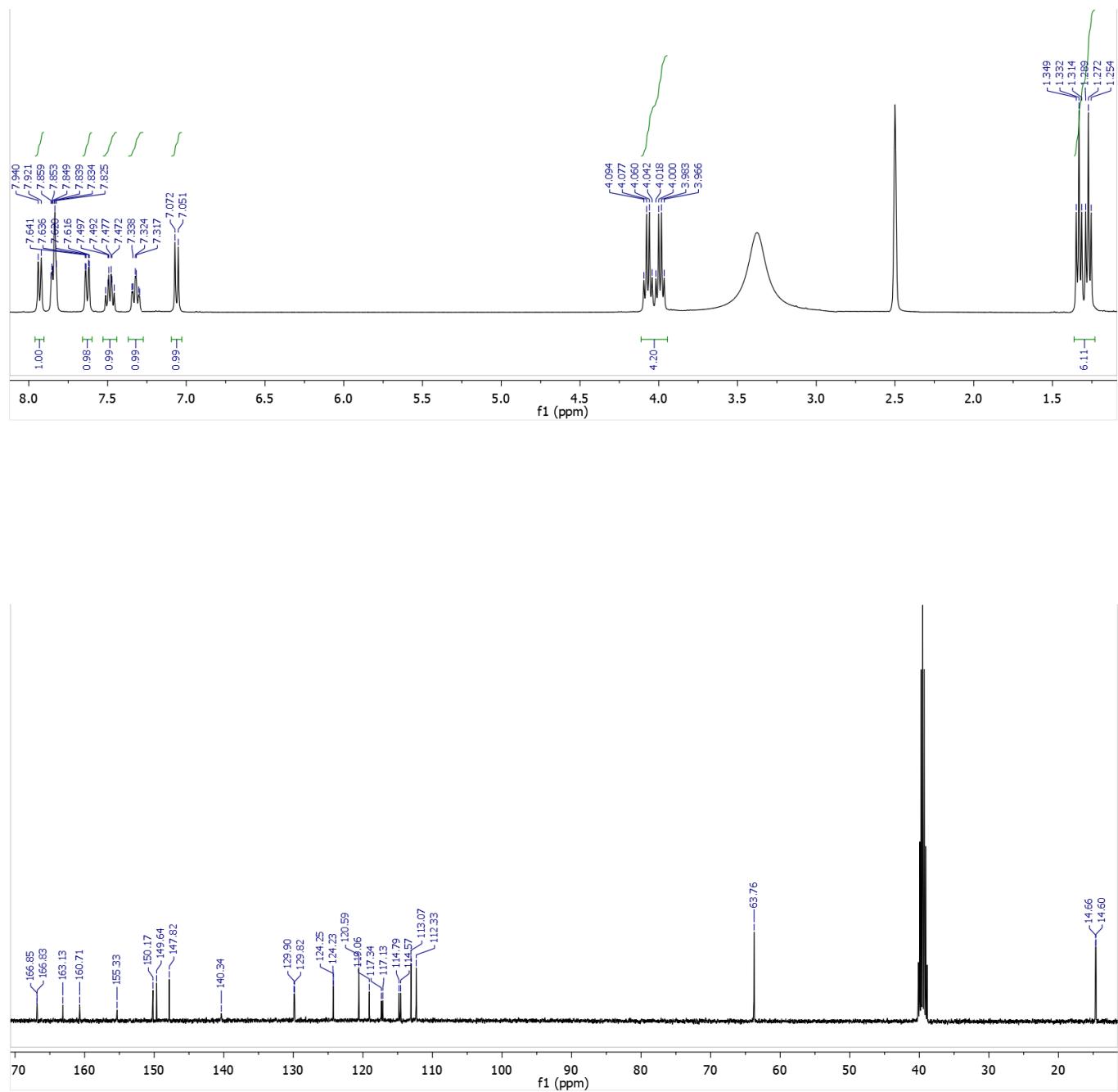


Figure S12. ^1H NMR at 400 MHz and ^{13}C NMR at 100 MHz spectra for compound 15

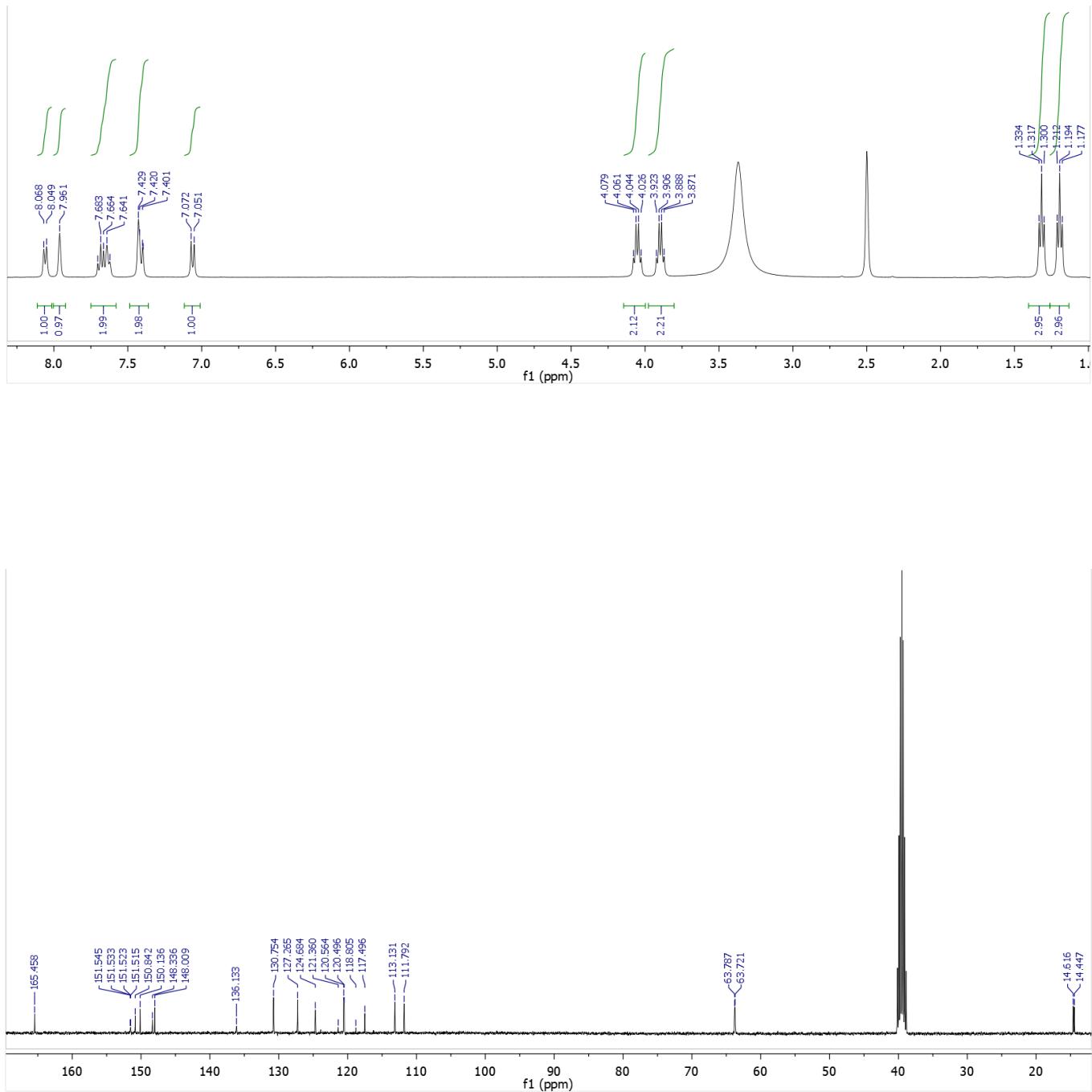


Figure S13. ^1H NMR at 400 MHz and ^{13}C NMR at 100 MHz spectra for compound 16

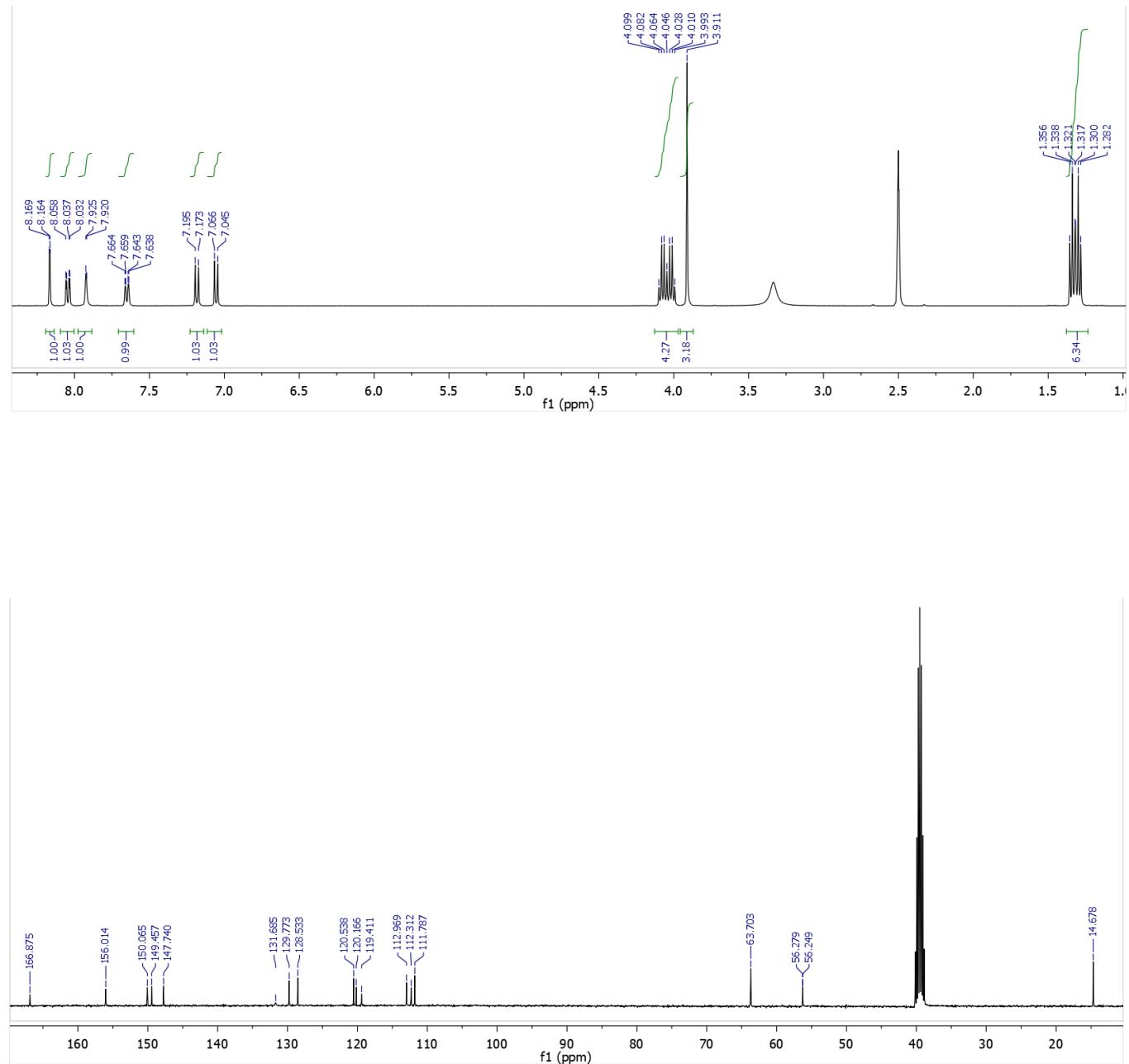


Figure S14. ^1H NMR at 400 MHz and ^{13}C NMR at 100 MHz spectra for compound 17

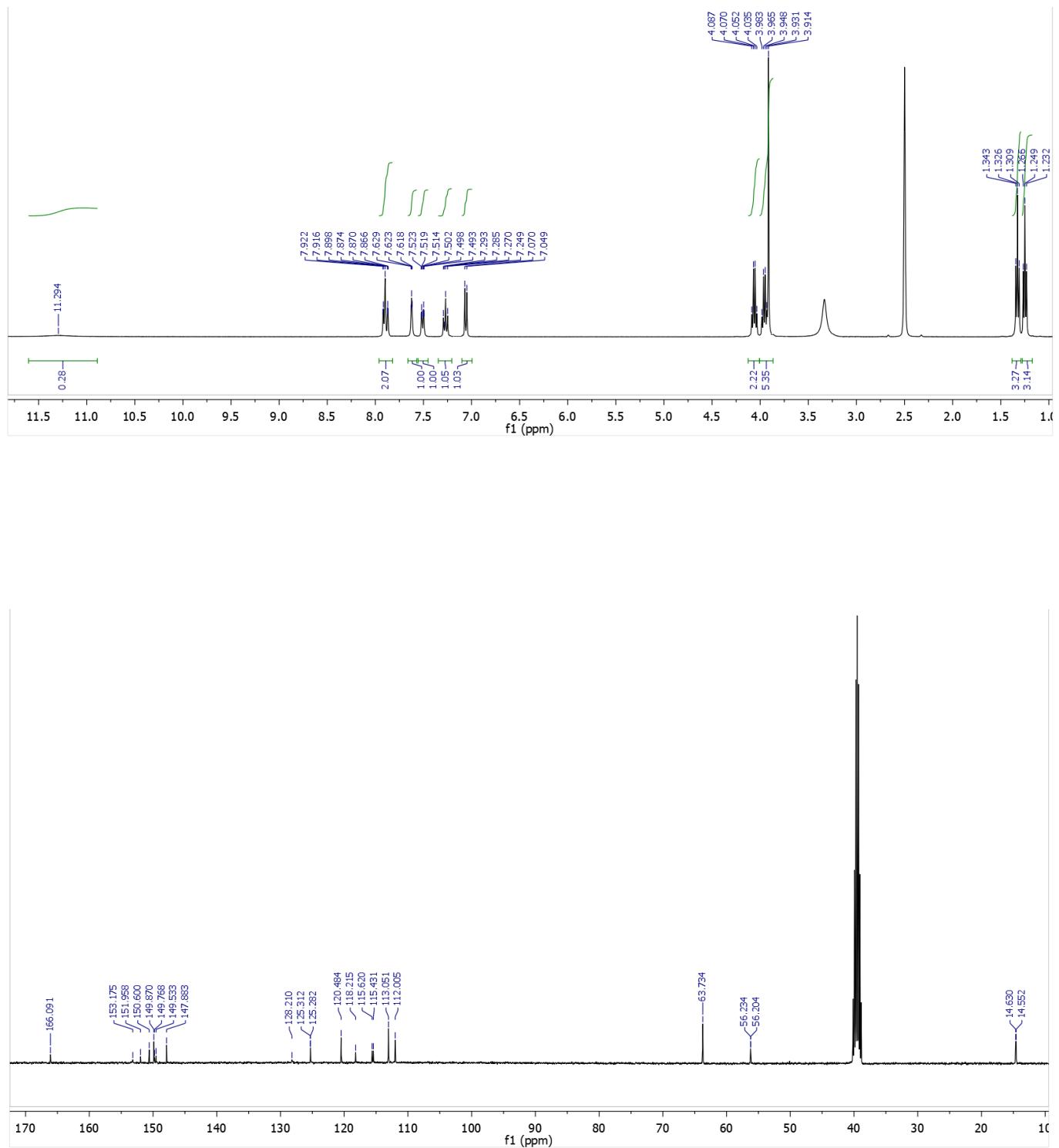


Figure S15. ^1H NMR at 400 MHz and ^{13}C NMR at 100 MHz spectra for compound 26

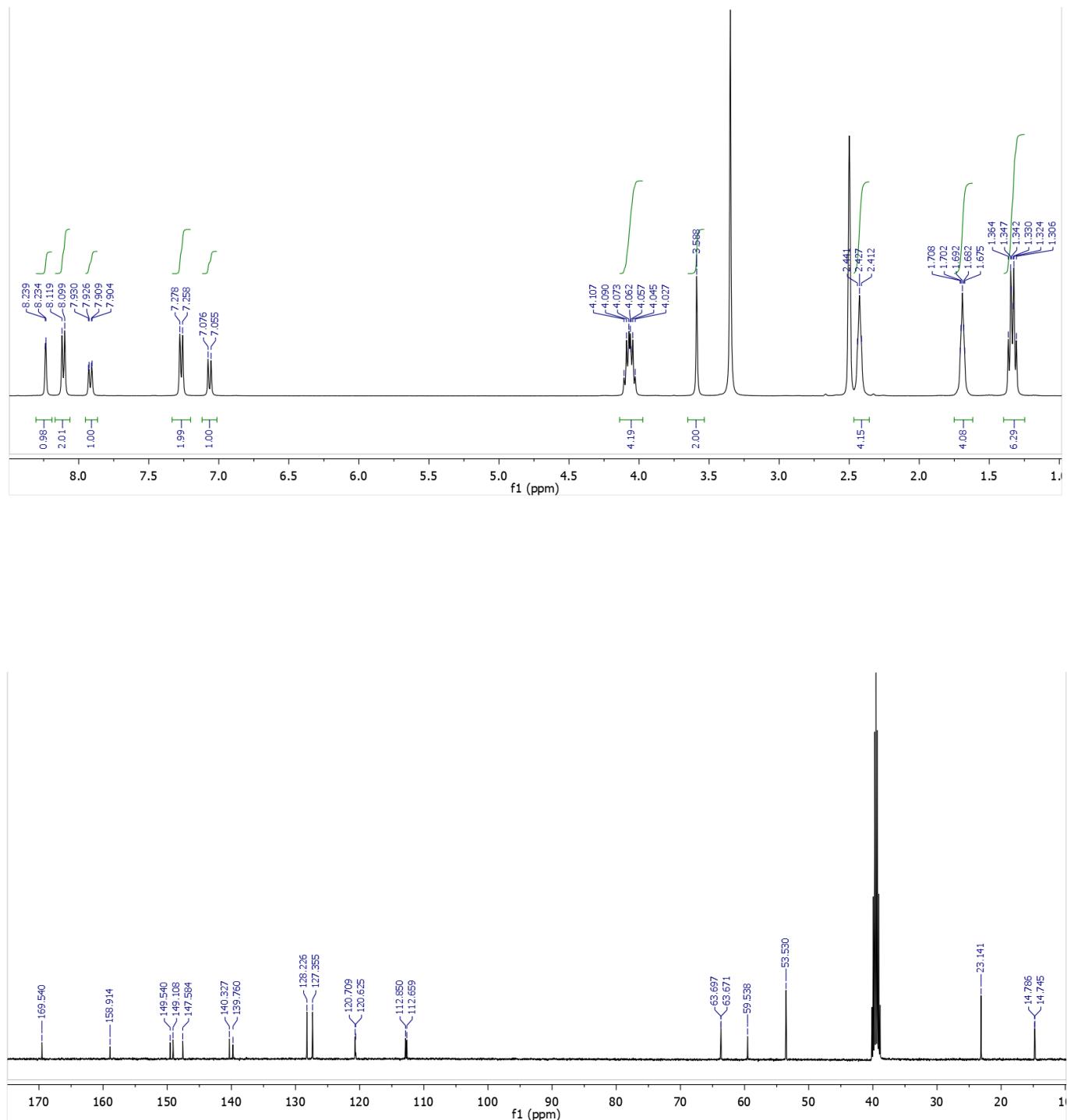


Figure S16. ^1H NMR at 400 MHz and ^{13}C NMR at 100 MHz spectra for compound 27

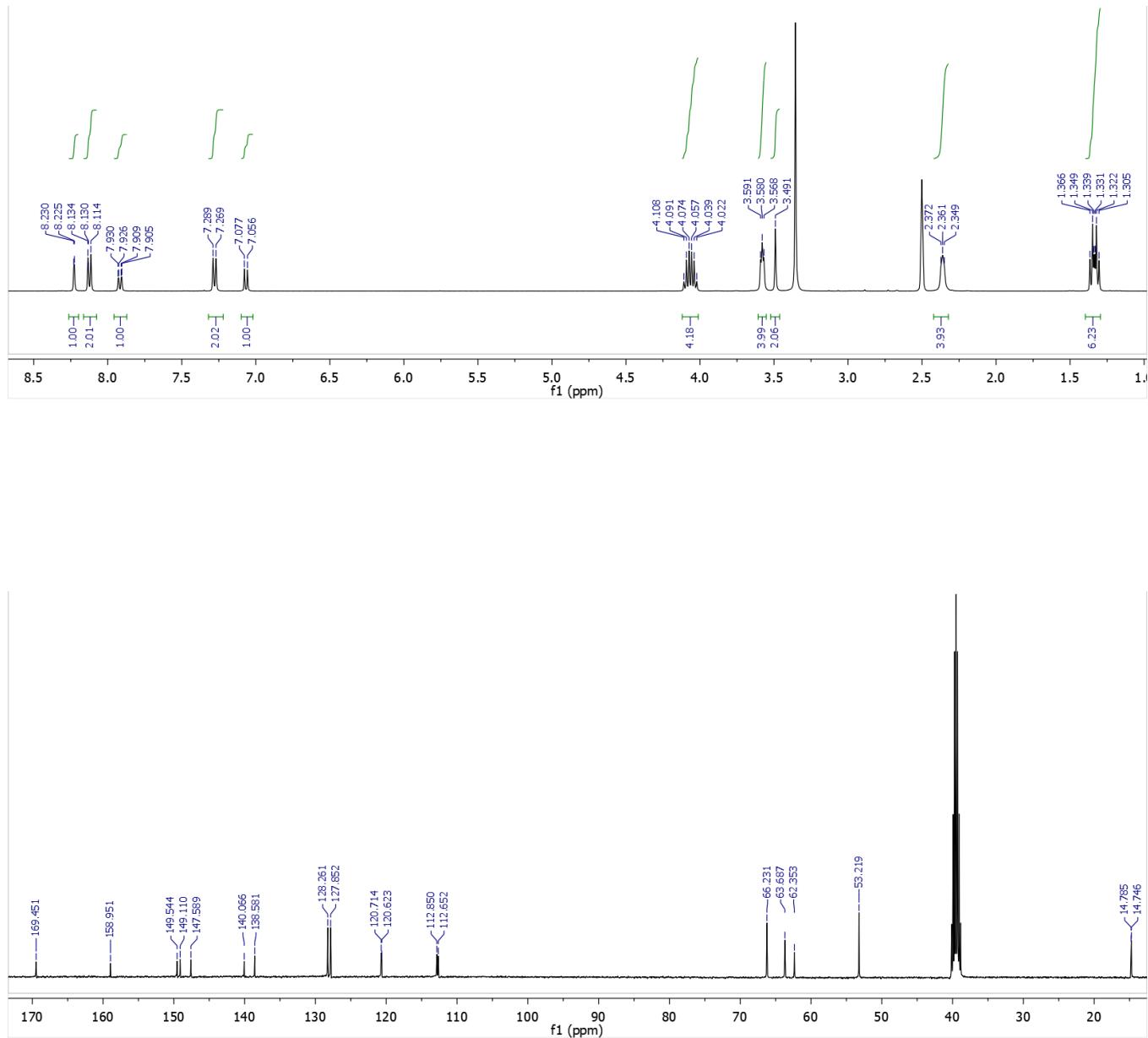


Figure S17. ^1H NMR at 400 MHz and ^{13}C NMR at 100 MHz spectra for compound 28

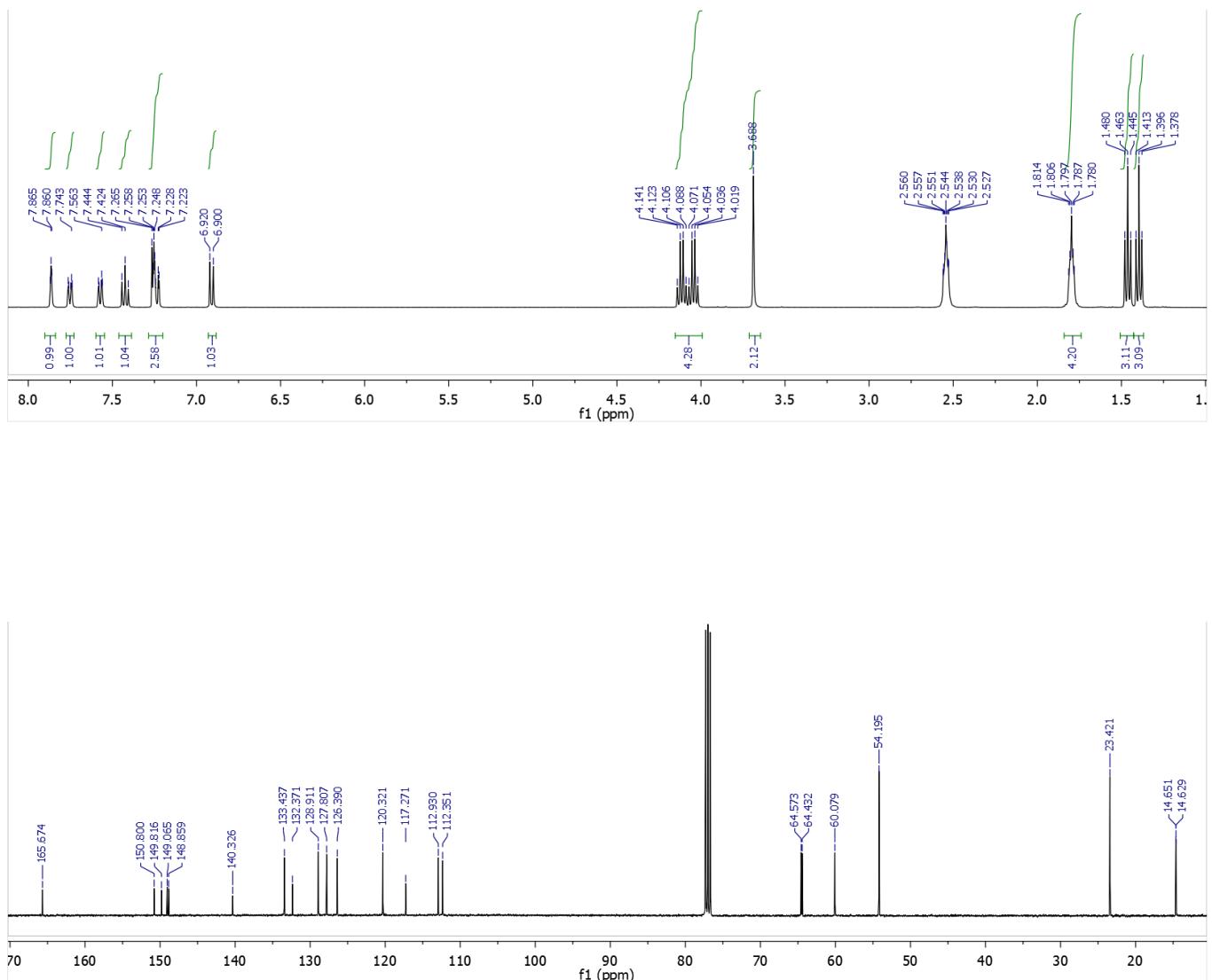


Figure S18. ^1H NMR at 400 MHz and ^{13}C NMR at 100 MHz spectra for compound **29**

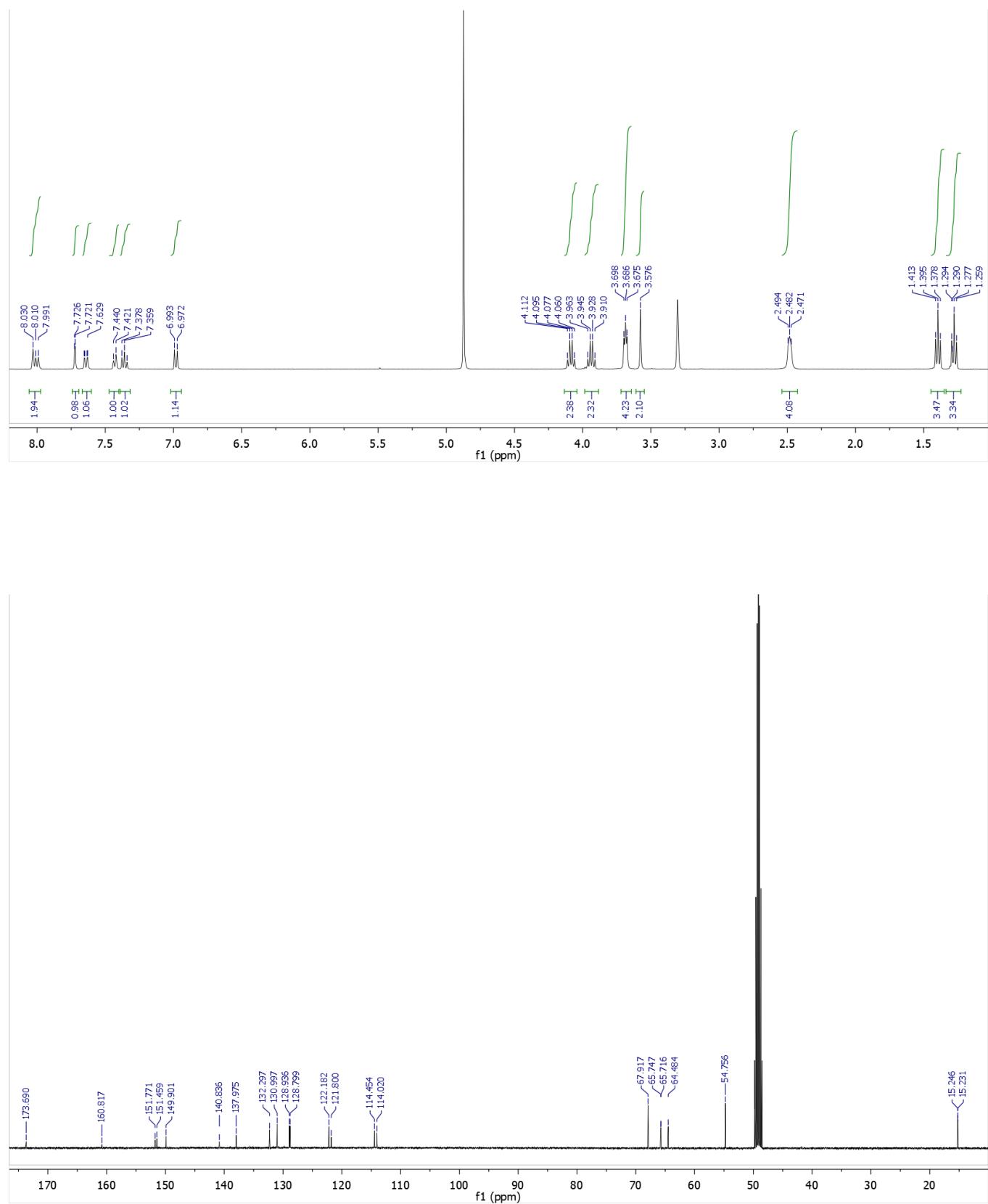


Figure S19. ^1H NMR at 400 MHz and ^{13}C NMR at 100 MHz spectra for compound 31

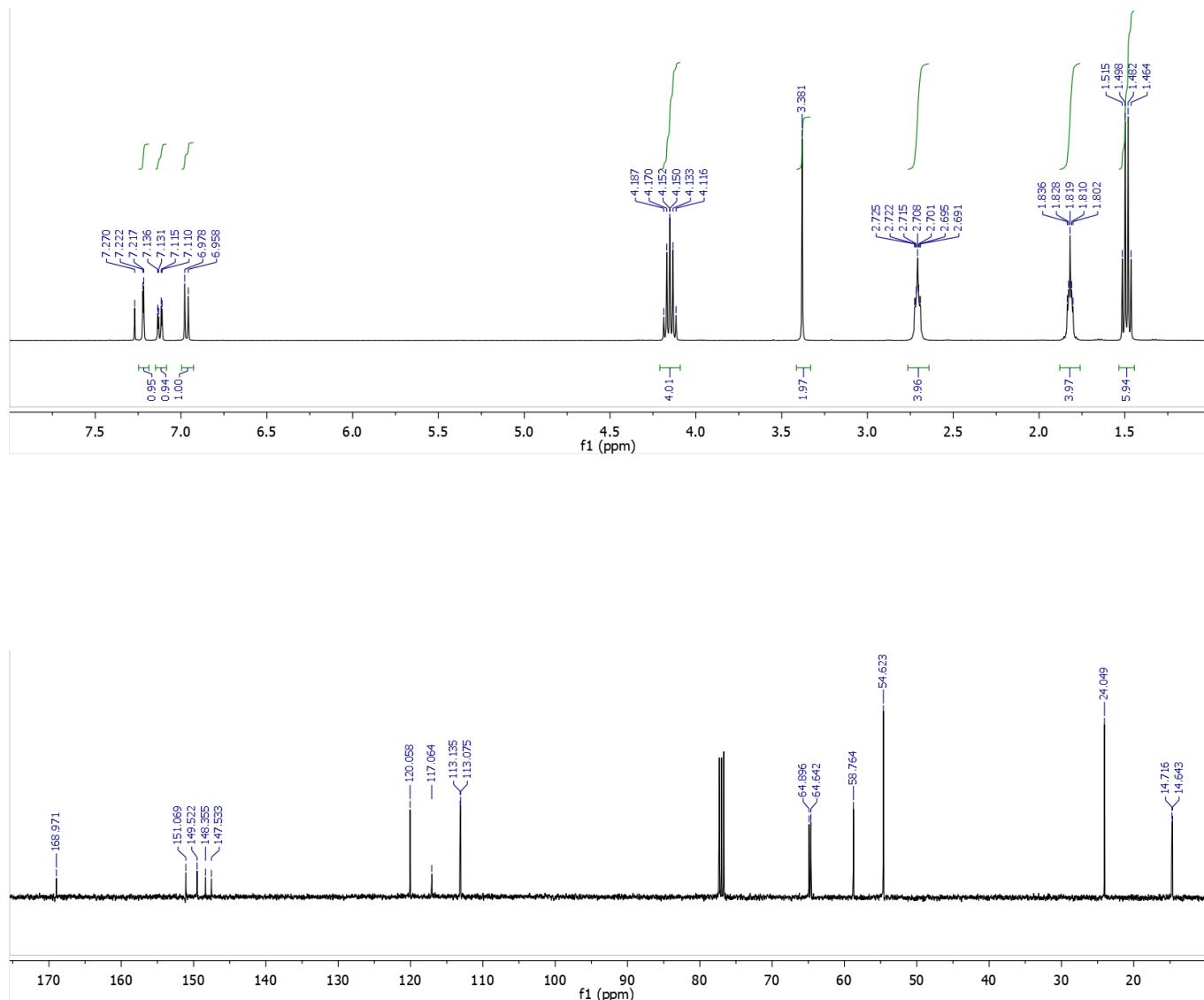


Figure S20. ^1H NMR at 400 MHz and ^{13}C NMR at 100 MHz spectra for compound 32

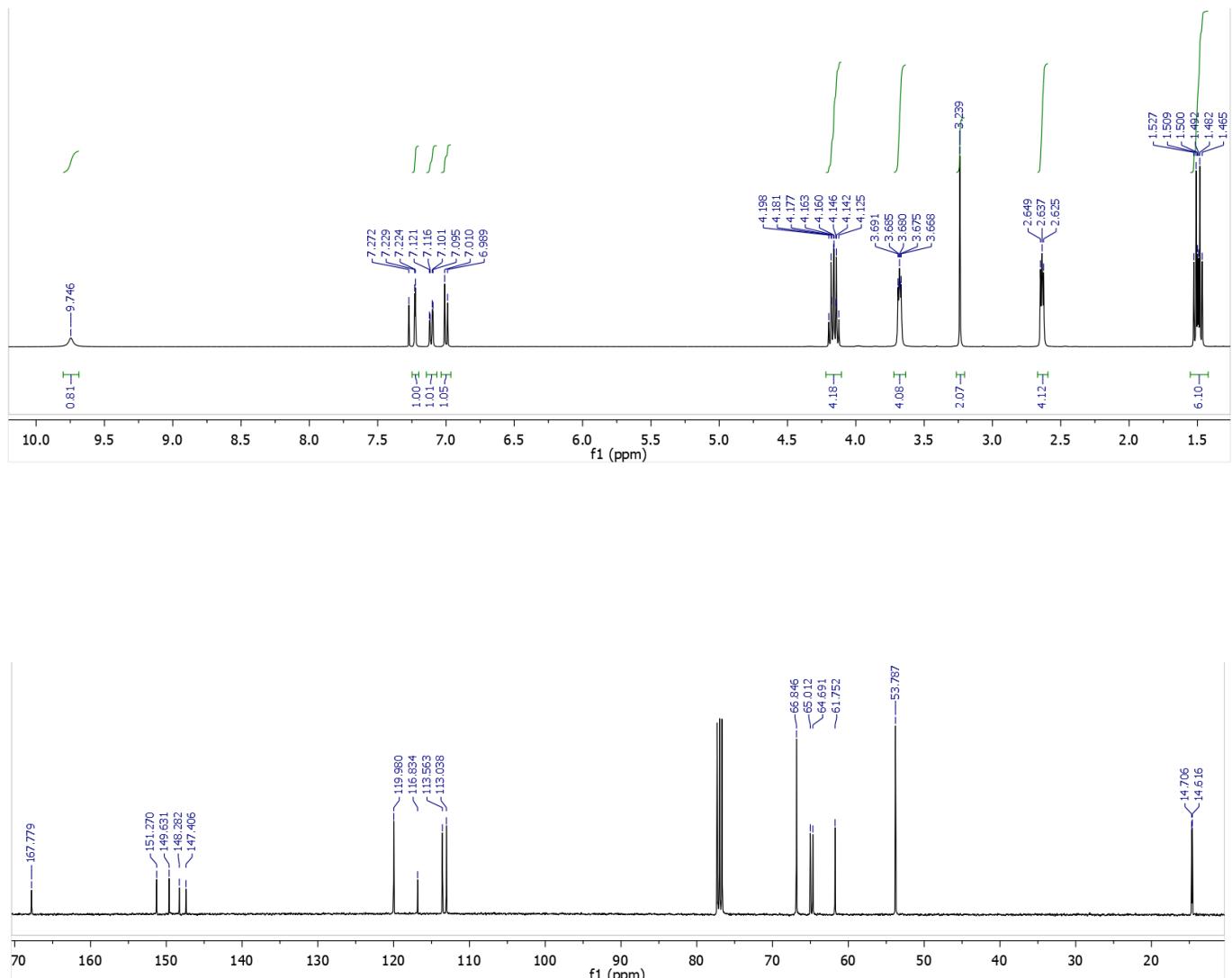


Figure S21. ^1H NMR at 400 MHz and ^{13}C NMR at 100 MHz spectra for compound 34

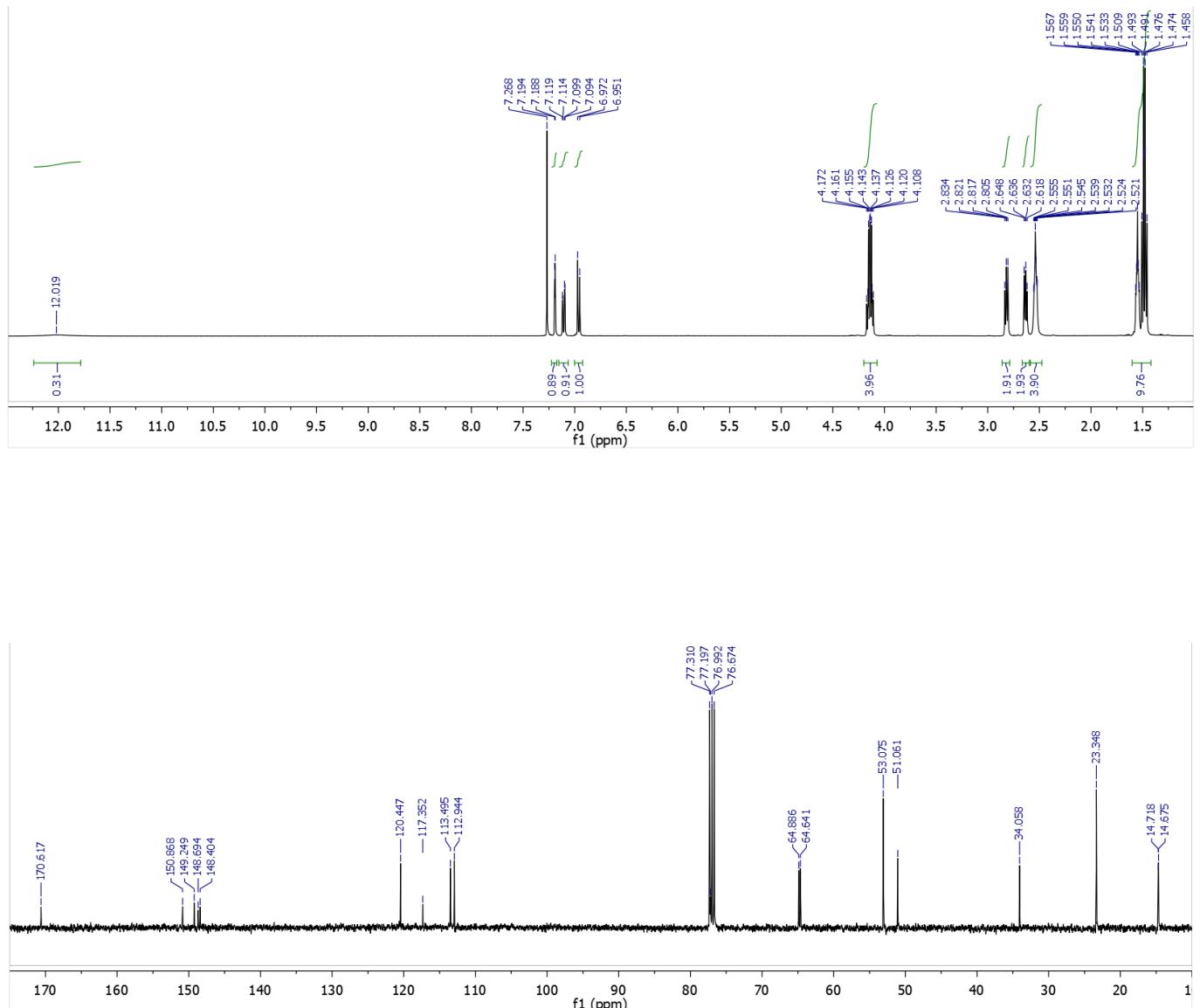


Figure S22. ^1H NMR at 400 MHz and ^{13}C NMR at 100 MHz spectra for compound 35

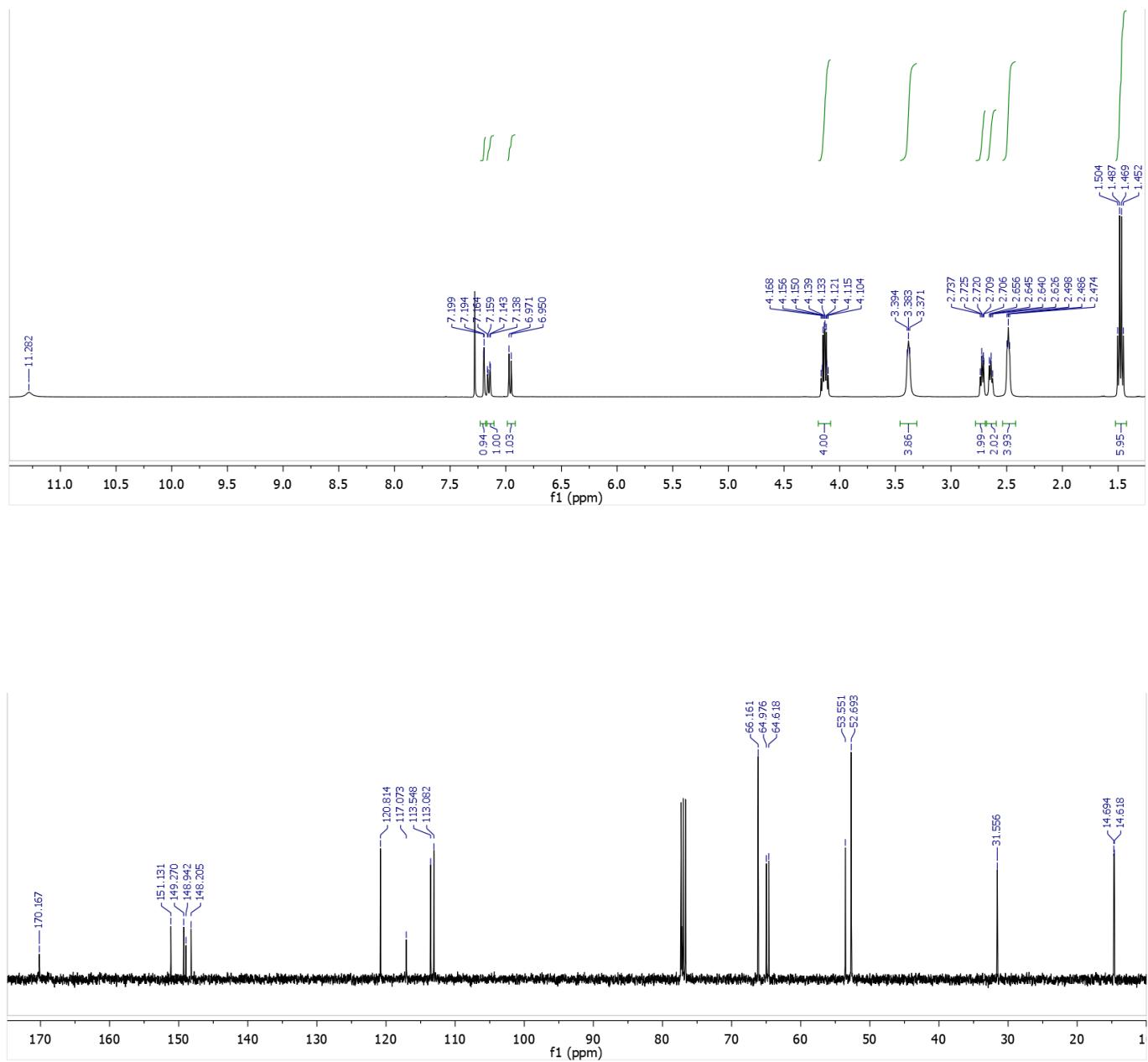


Figure S23. ^1H NMR at 400 MHz and ^{13}C NMR at 100 MHz spectra for compound 39

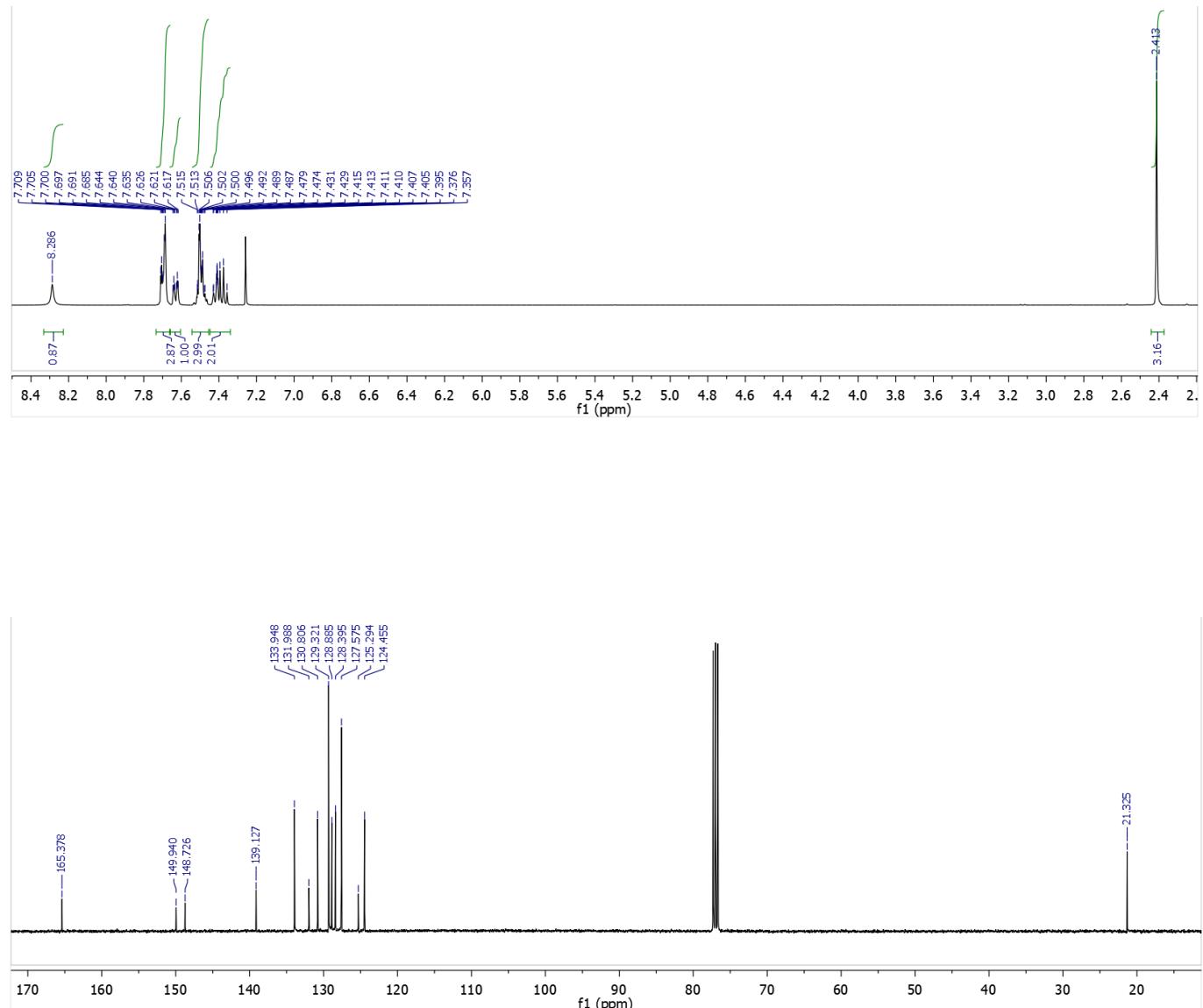


Figure S24. Dose-response curves of compound **1** against *P.f.* NF54, *P.f.* K₁ and L-6 cells.

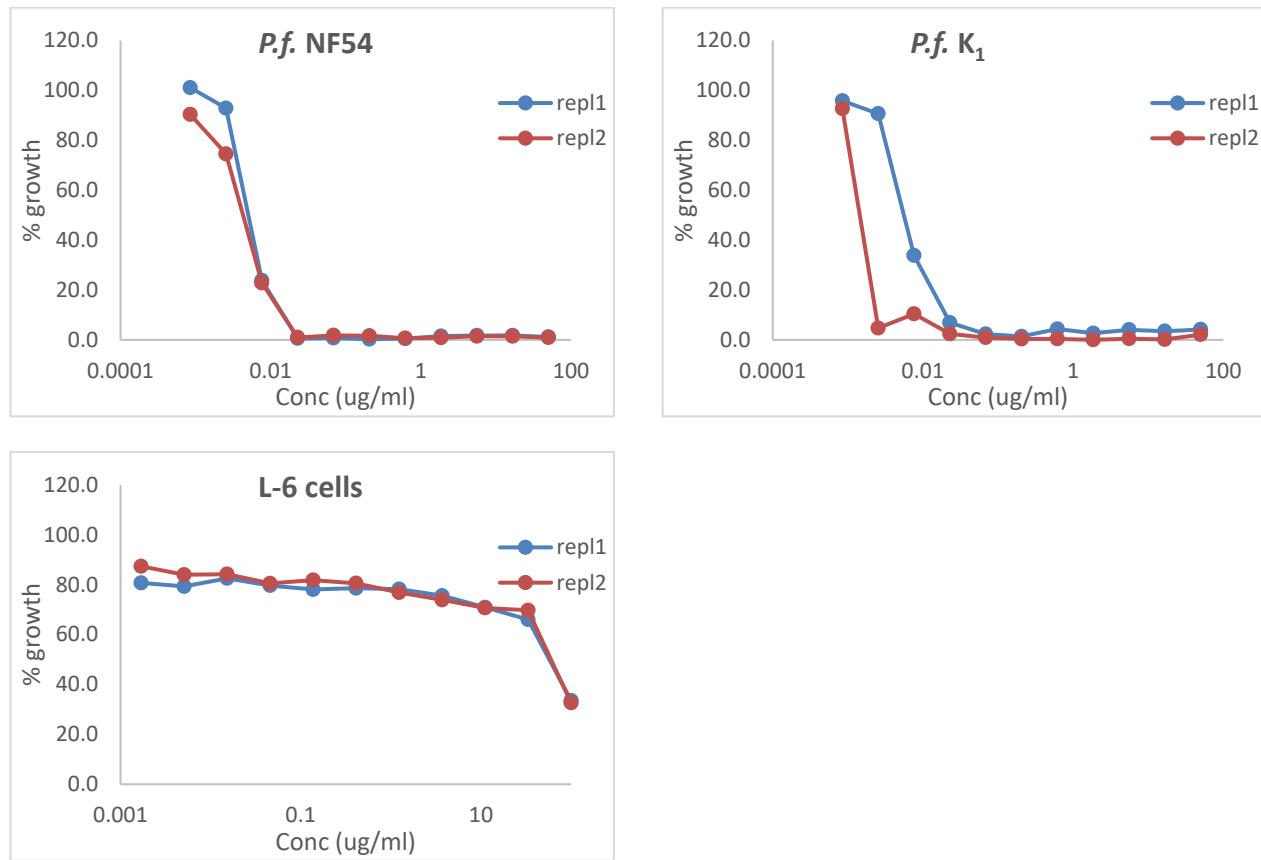


Figure S25. Dose-response curves of compound **10** against *P.f.* NF54, *P.f.* K₁ and L-6 cells.

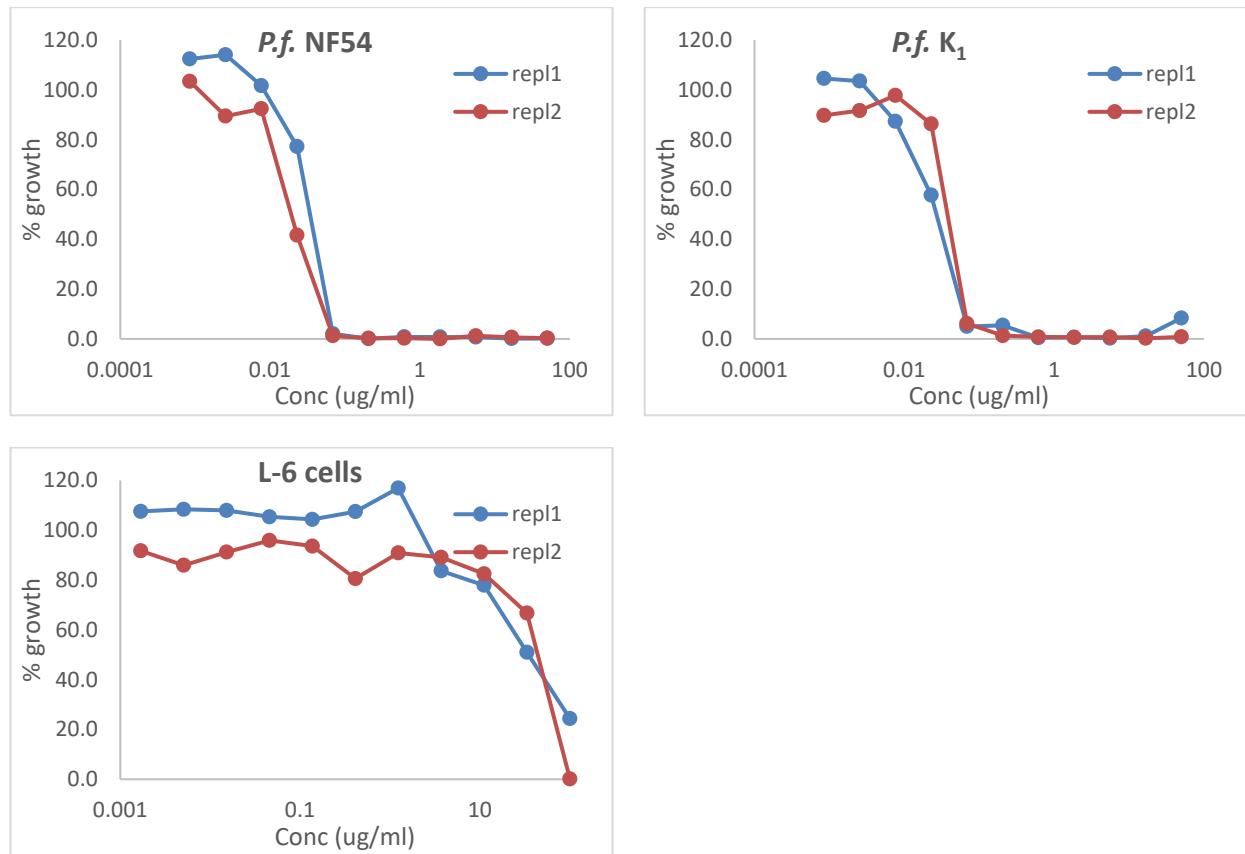


Figure S26. Dose-response curves of compound **11** against *P.f.* NF54 and L-6 cells.

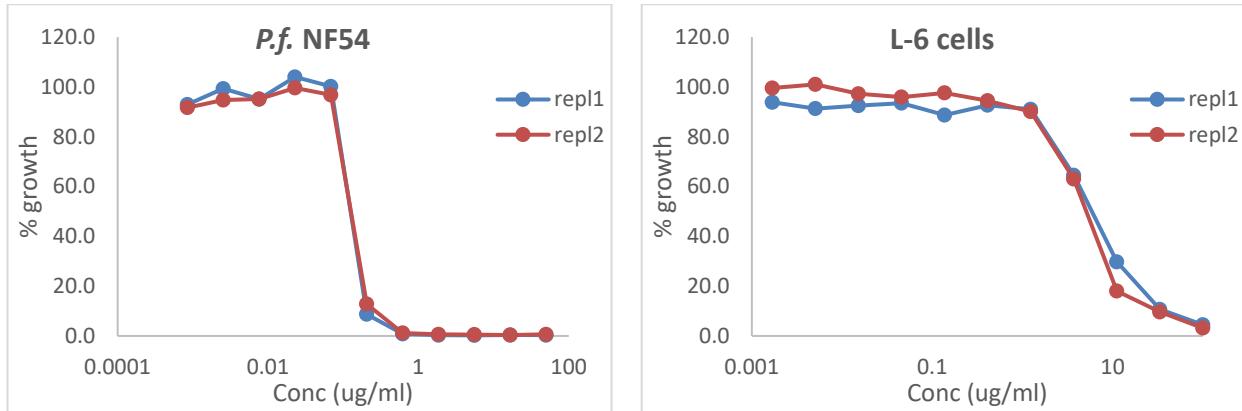


Figure S27. Dose-response curves of compound **12** against *P.f.* NF54 and L-6 cells.

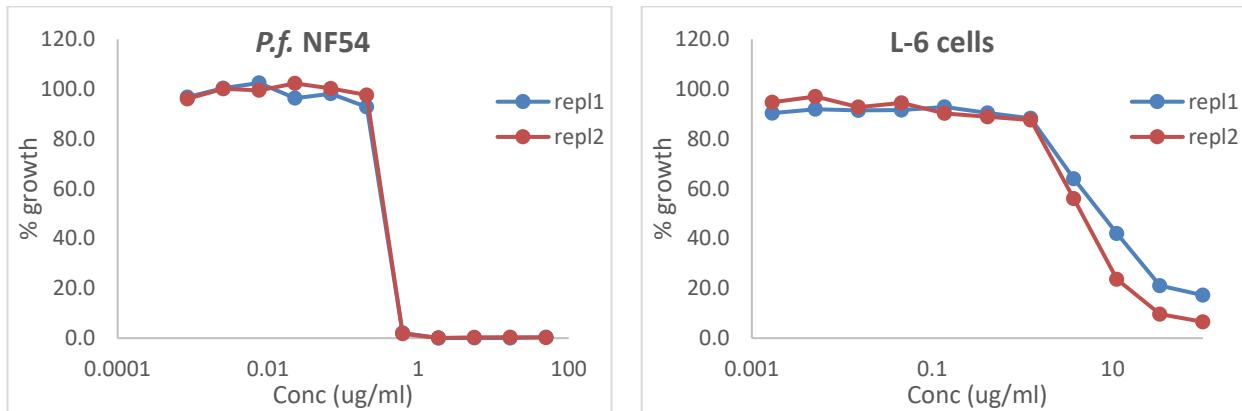
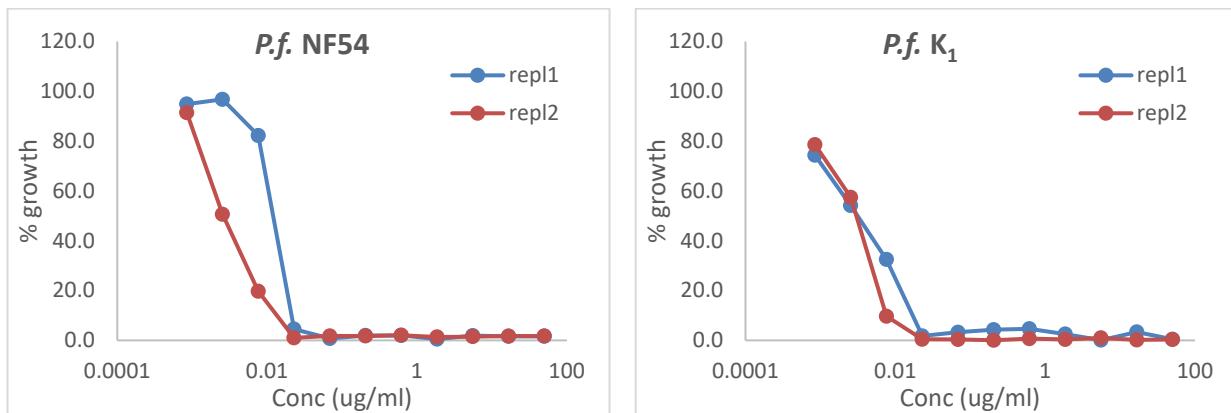


Figure S28. Dose-response curves of compound **13** against *P.f.* NF54, *P.f.* K₁ and L-6 cells.



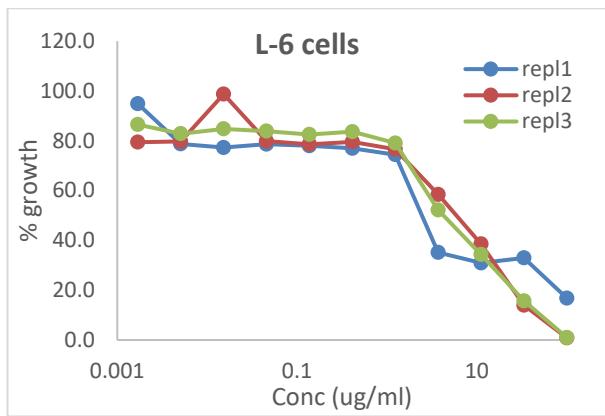


Figure S29. Dose-response curves of compound **14** against *P.f.* NF54, *P.f.* K₁ and L-6 cells.

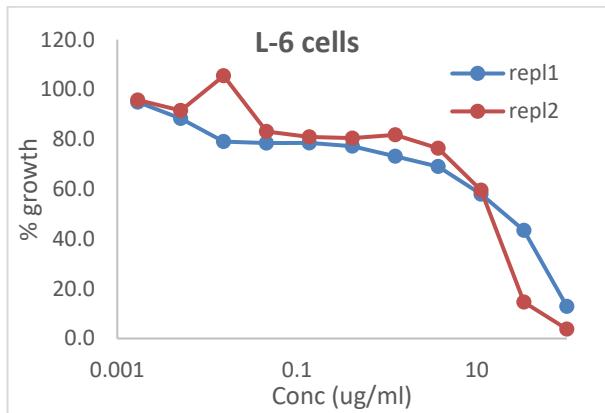
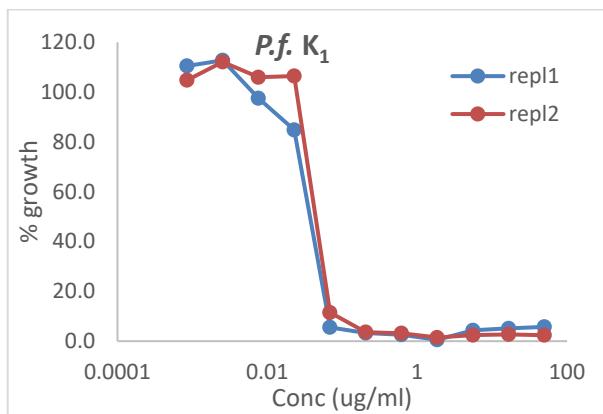
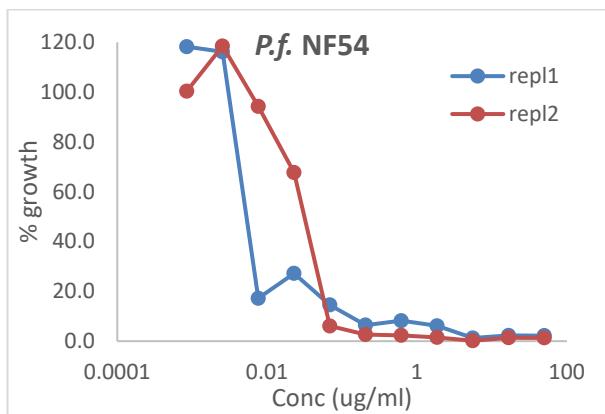


Figure S30. Dose-response curves of compound **15** against *P.f.* NF54 and L-6 cells.

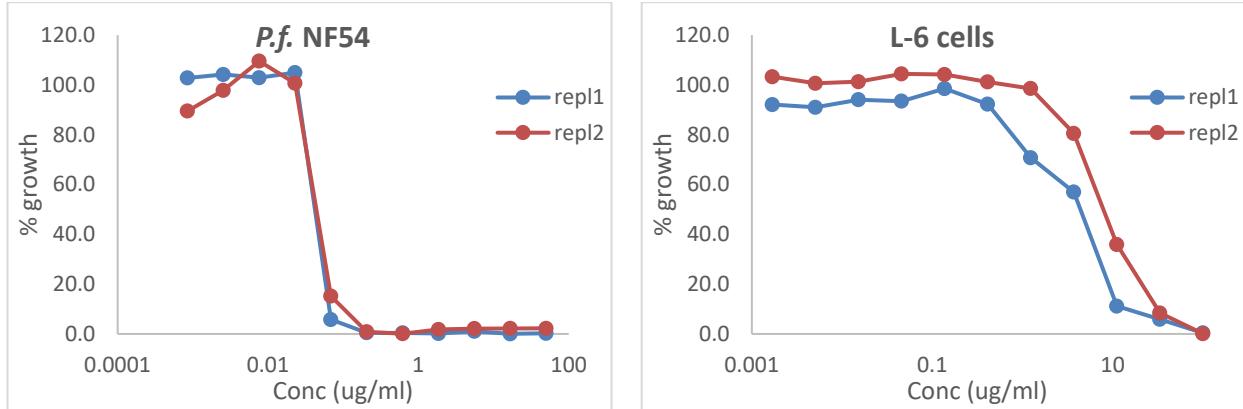


Figure S31. Dose-response curves of compound **16** against *P.f.* NF54, *P.f.* K₁ and L-6 cells.

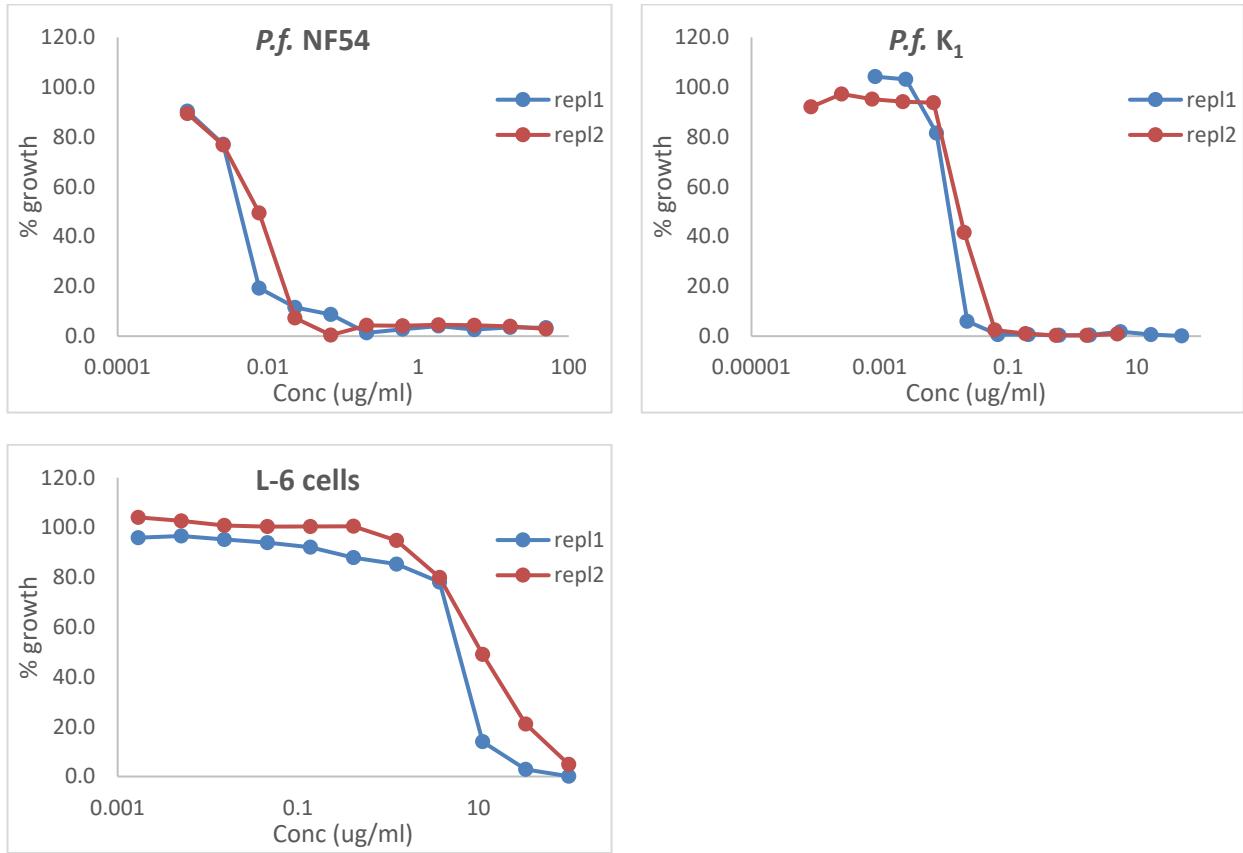


Figure S32. Dose-response curves of compound 17 against *P.f.* NF54, *P.f.* K₁ and L-6 cells.

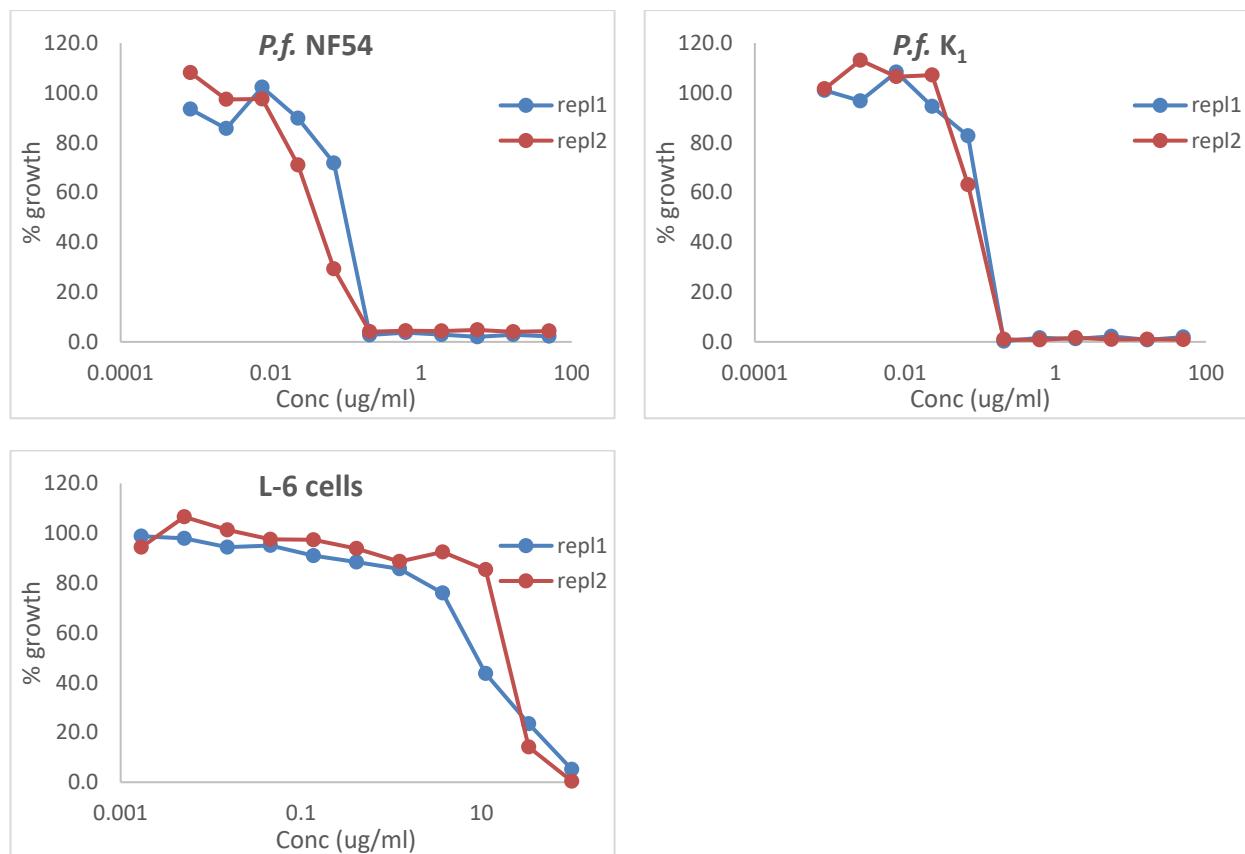
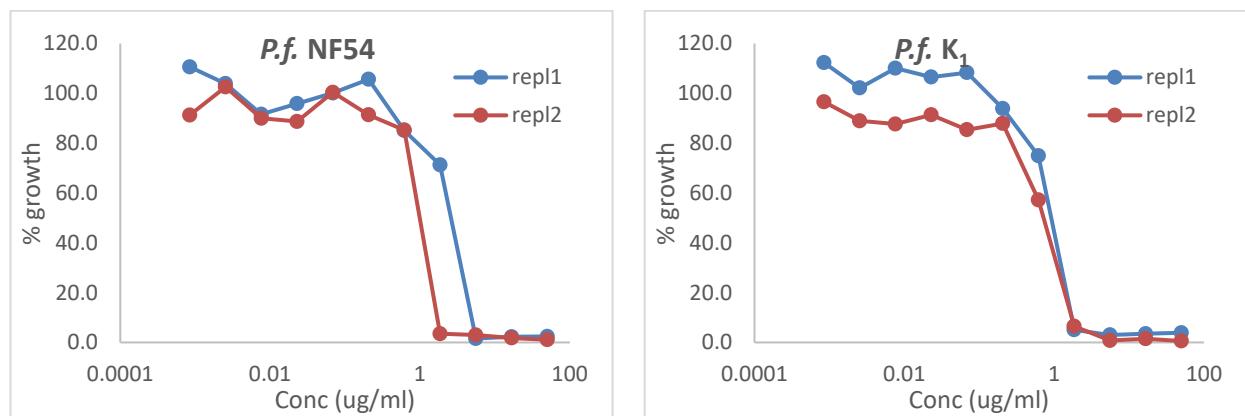


Figure S33. Dose-response curves of compound 26 against *P.f.* NF54, *P.f.* K₁ and L-6 cells.



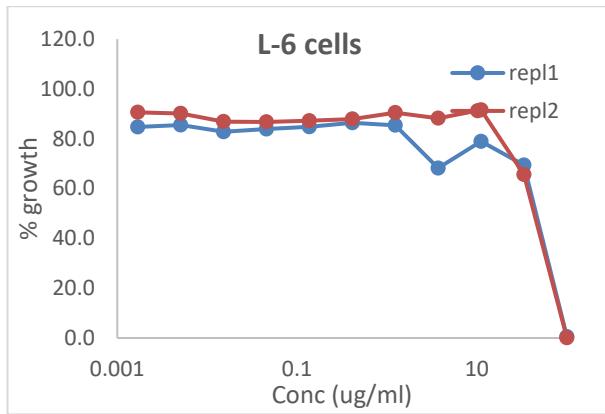


Figure S34. Dose-response curves of compound **27** against *P.f.* NF54, *P.f.* K₁ and L-6 cells.

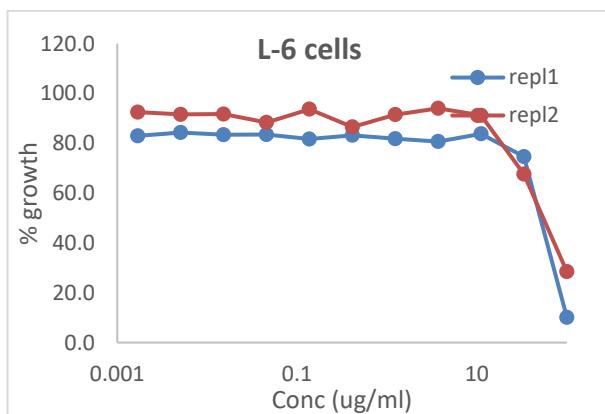
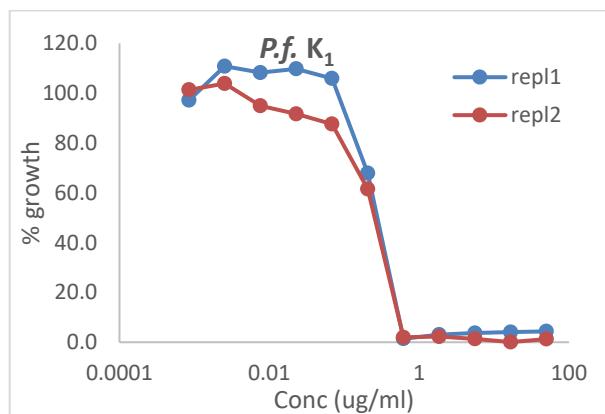
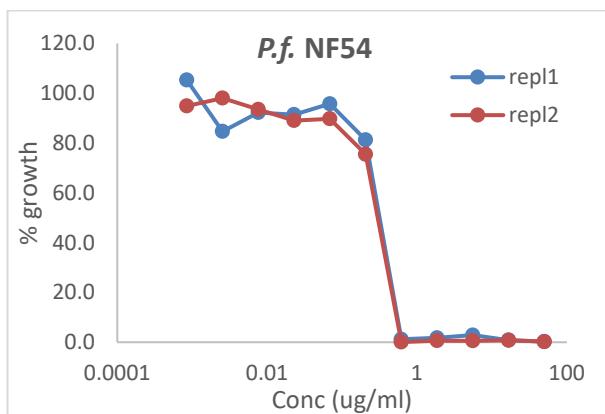


Figure S35. Dose-response curves of compound **28** against *P.f.* NF54 and L-6 cells.

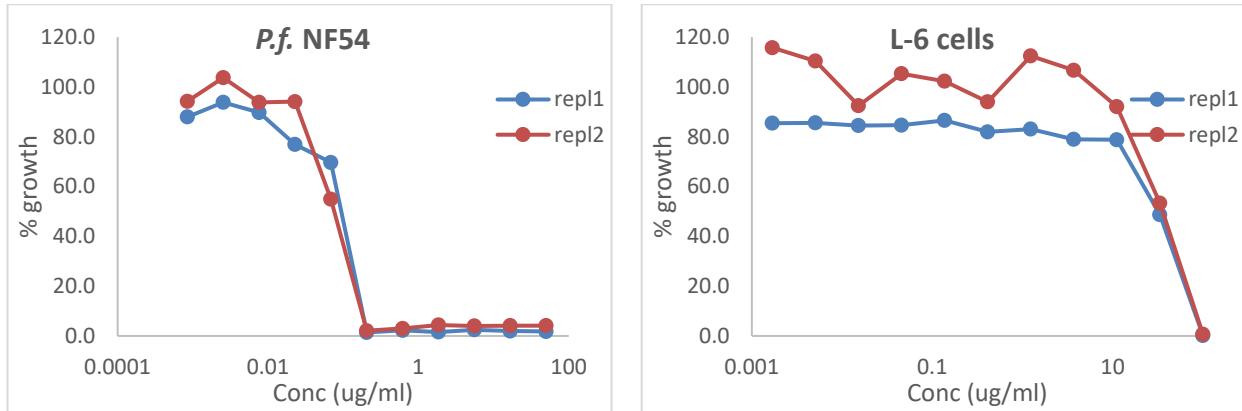


Figure S36. Dose-response curves of compound **29** against *P.f.* NF54 and L-6 cells.

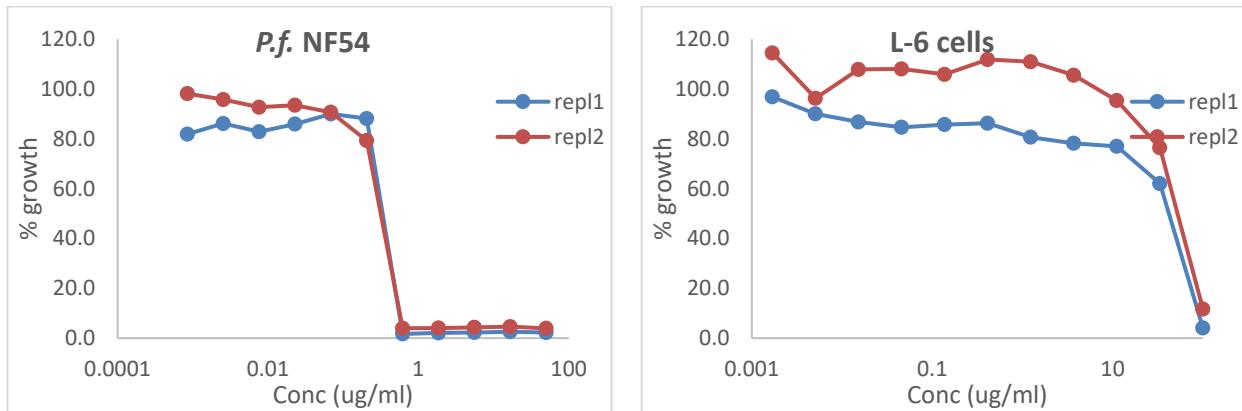
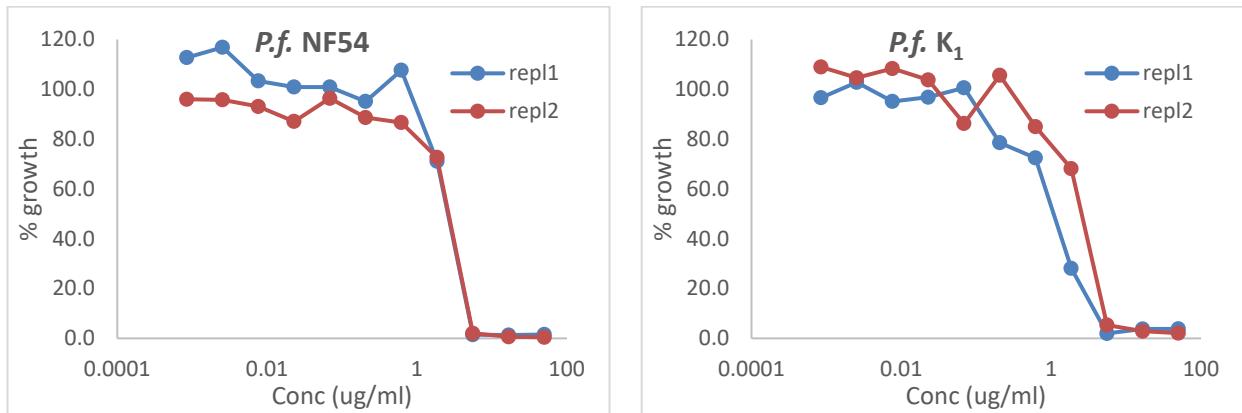


Figure S37. Dose-response curves of compound **31** against *P.f.* NF54, *P.f.* K₁ and L-6 cells.



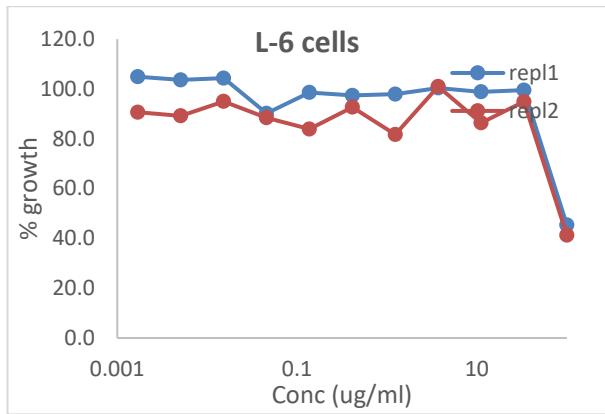


Figure S38. Dose-response curves of compound **32** against *P.f.* NF54, *P.f.* K₁ and L-6 cells.

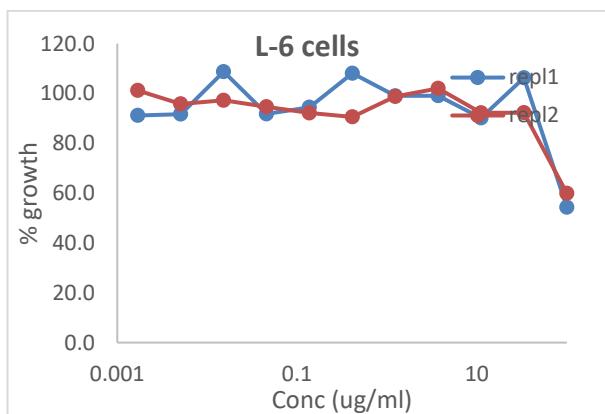
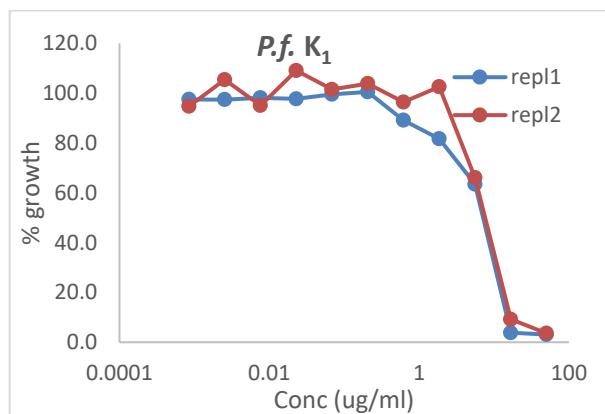
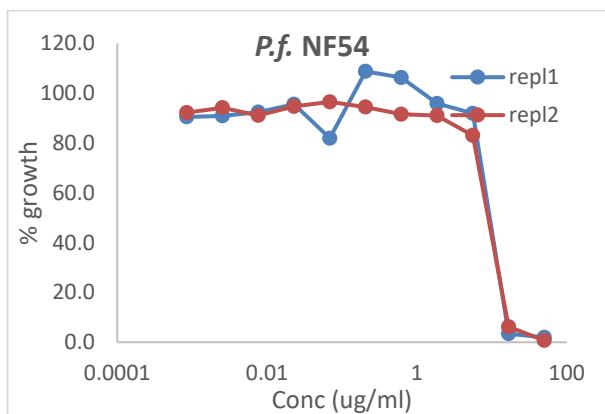


Figure S39. Dose-response curves of compound **34** against *P.f.* NF54, *P.f.* K₁ and L-6 cells.

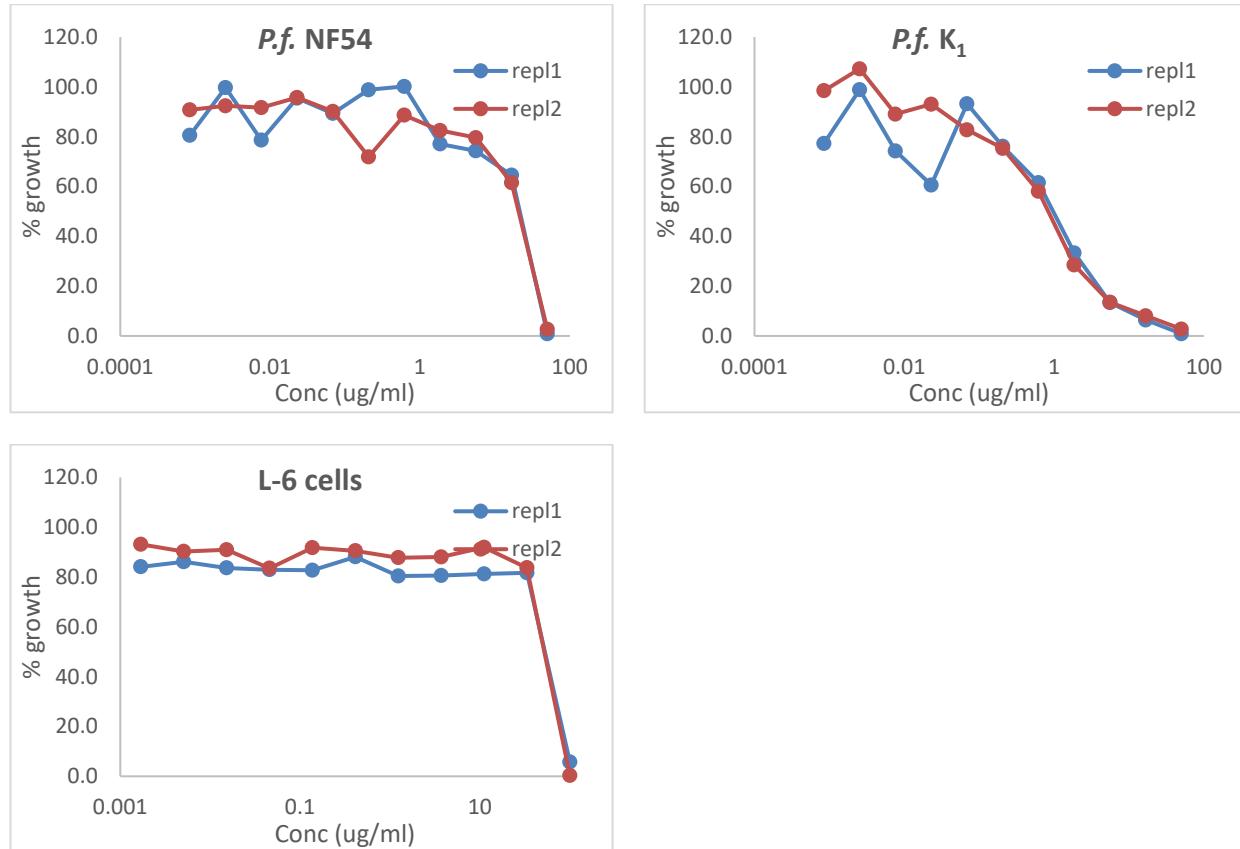


Figure S40. Dose-response curves of compound **35** against *P.f.* NF54, *P.f.* K₁ and L-6 cells.

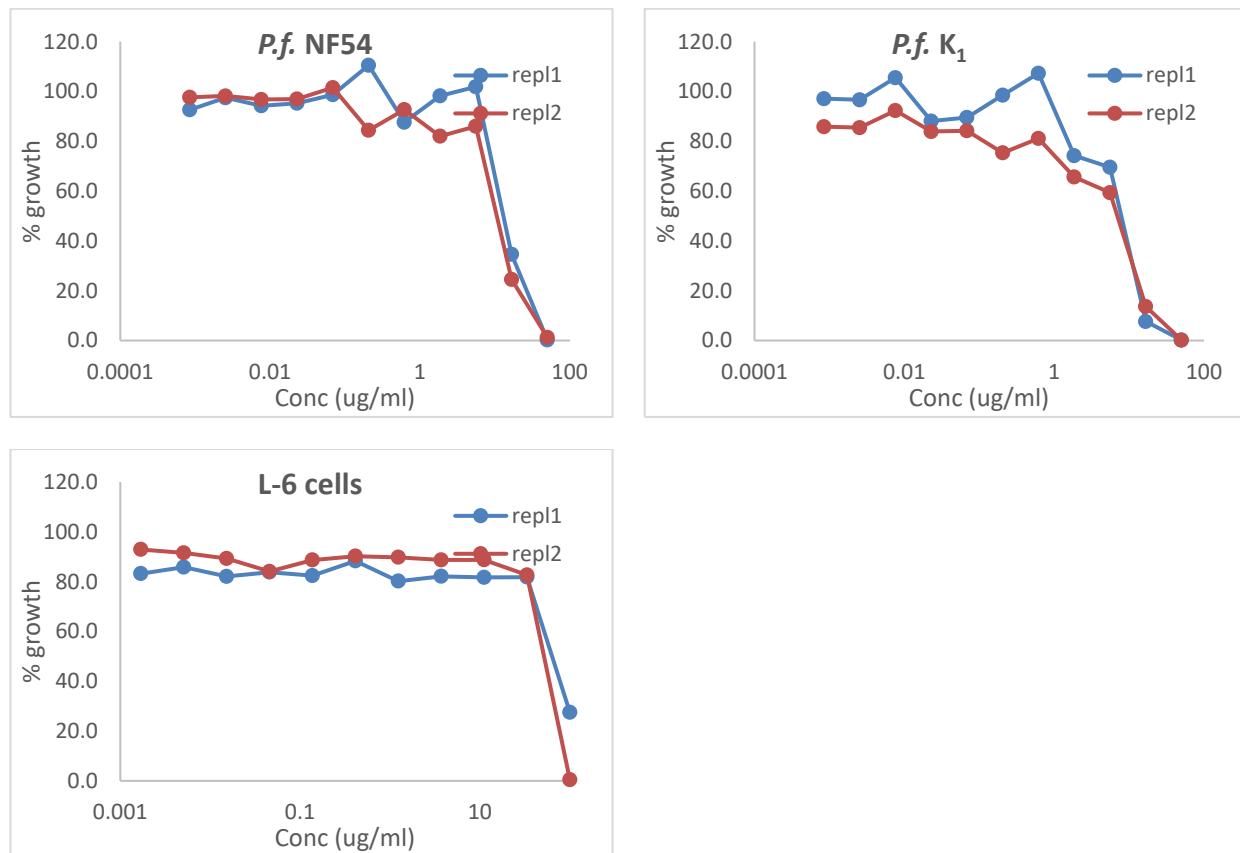


Figure S41. Dose-response curves of compound **39** against *P.f.* NF54, *P.f.* K₁ and L-6 cells.

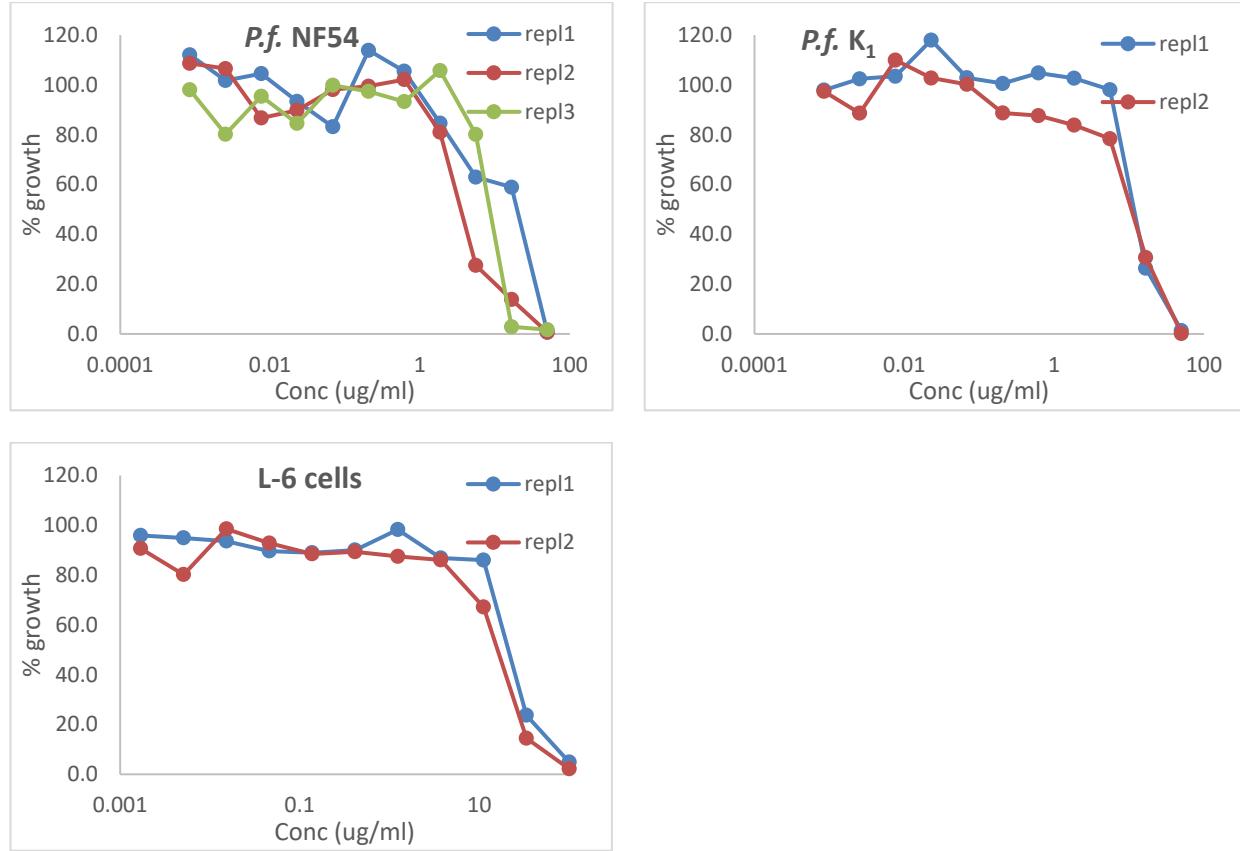


Table S1. Activities with standard deviation values of compounds **1**, **10-17**, **26-29**, **31**, **32**, **34**, **35** and **39** against *P.falciparum* NF54, *P.falciparum* K₁ and L-6 cells, expressed as IC₅₀ (μM).^a

Compound	<i>P.f.</i> NF54 ^b	<i>P.f.</i> K ₁ ^c	Cytotoxicity L-6 cells
1	0.011 ± 0.001	0.011 ± 0.0068	159.3 ± 3.36
10	0.076 ± 0.021	0.091 ± 0.015	111.2 ± 12.9
11	0.343 ± 0.001		14.92 ± 1.05
12	0.831 ± 0.011		14.24 ± 3.45
13	0.019 ± 0.010	0.007 ± 0.00	9.97 ± 3.36
14	0.049 ± 0.035	0.108 ± 0.025	41.09 ± 9.96
15	0.098 ± 0.002		13.99 ± 3.97
16	0.014 ± 0.003		20.03 ± 5.67
17	0.167 ± 0.068		35.03 ± 12.7
26	4.055 ± 1.78	1.87 ± 0.210	101.7 ± 2.41
27	0.674 ± 0.015	0.586 ± 0.020	116.4 ± 4.35
28	0.192 ± 0.020		77.23 ± 4.46
29	0.712 ± 0.013		104.0 ± 11.4
31	7.24 ± 0.068	5.02 ± 2.03	242.3 ± 10.3
32	24.25 ± 0.575	19.53 ± 0.610	265.7 ± 0.00
34	56.01 ± 0.821	2.41 ± 0.179	139.8 ± 0.995
35	30.15 ± 3.05	18.99 ± 1.15	147.3 ± 15.1
39	21.52 ± 8.99	39.92 ± 1.64	65.95 ± 9.06
ART		0.0064	450.5
CQ		0.15	188.5
POD			0.012

ART = artemisinin; CQ = chloroquine; POD = podophyllotoxin

^a Values represent the average of four determinations (two determinations of two independent experiments)

^b Sensitive to chloroquine

^c Resistant to chloroquine and pyrimethamine.