

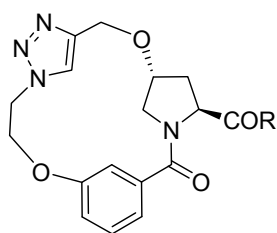
Supplementary Materials: Synthetic Strategy and Anti-Tumor Activities of Macrocyclic Scaffolds Based on 4-Hydroxyproline

Guorui Cao, Kun Yang, Yue Li, Longjiang Huang and Dawei Teng

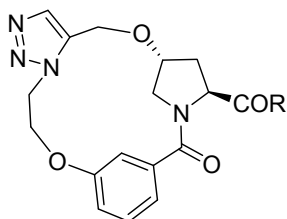
Table of Contents

1. General	S2
2. Spectra of macrocycle 6a–b	S3–S4
3. Spectra of macrocycle 9	S5
4. Spectra of macrocycle 10a–c	S6–S8
5. Spectra of macrocycle 22–45	S9–S20

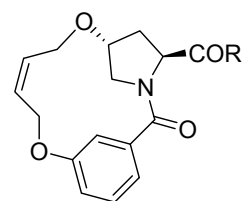
General



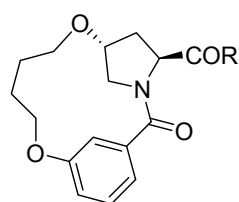
- 6a** R = OCH₃
22 R = OH
23 R = NH₂
24 R = NHPh
25 R = NHCH₂CH(CH₃)₂



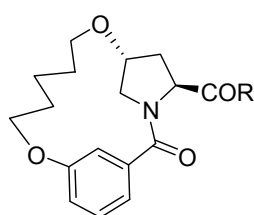
- 6b** R = OCH₃
26 R = OH
27 R = NH₂
28 R = NHPh
29 R = NHCH₂CH(CH₃)₂



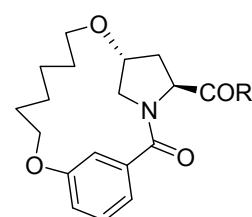
- 9** R = OCH₃
30 R = OH
31 R = NH₂
32 R = NHPh
33 R = NHCH₂CH(CH₃)₂



- 10a** R = OCH₃
34 R = OH
35 R = NH₂
36 R = NHPh
37 R = NHCH₂CH(CH₃)₂



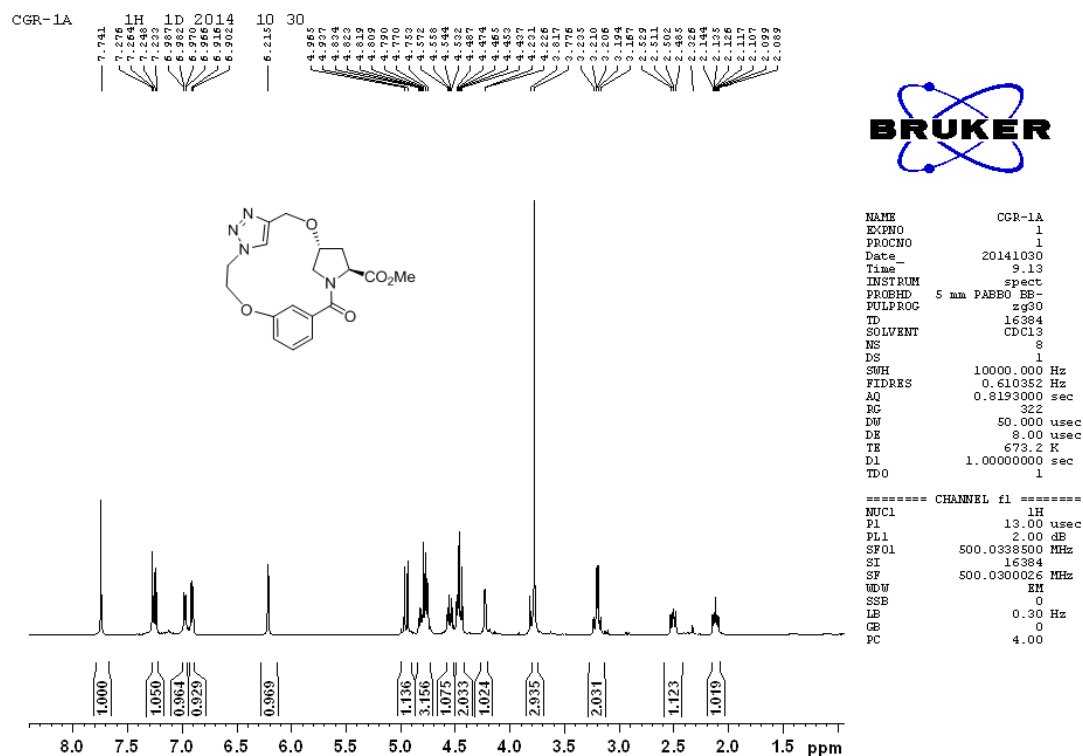
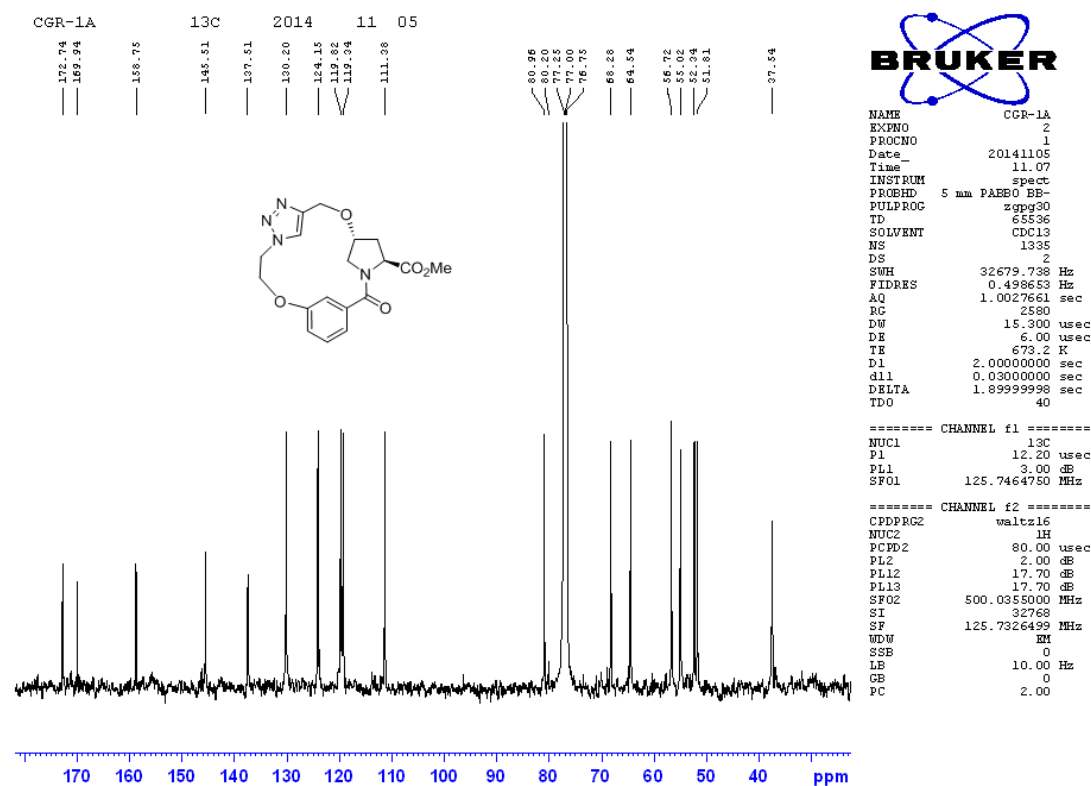
- 10b** R = OCH₃
38 R = OH
39 R = NH₂
40 R = NHPh
41 R = NHCH₂CH(CH₃)₂

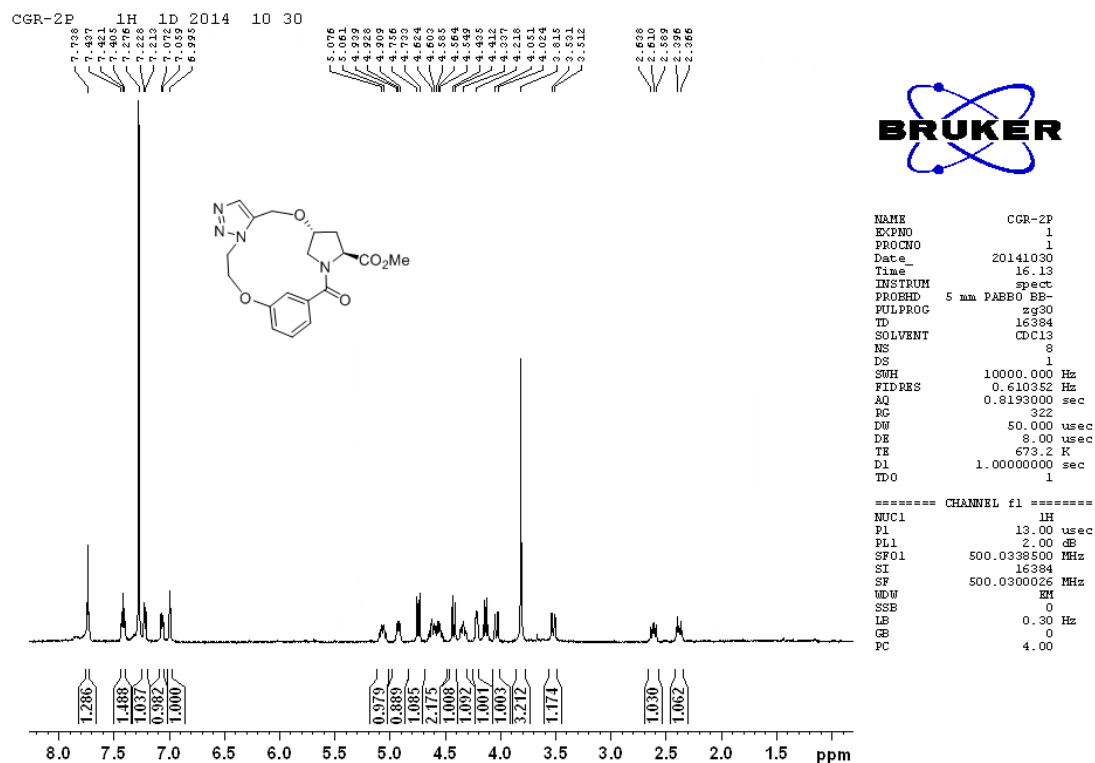
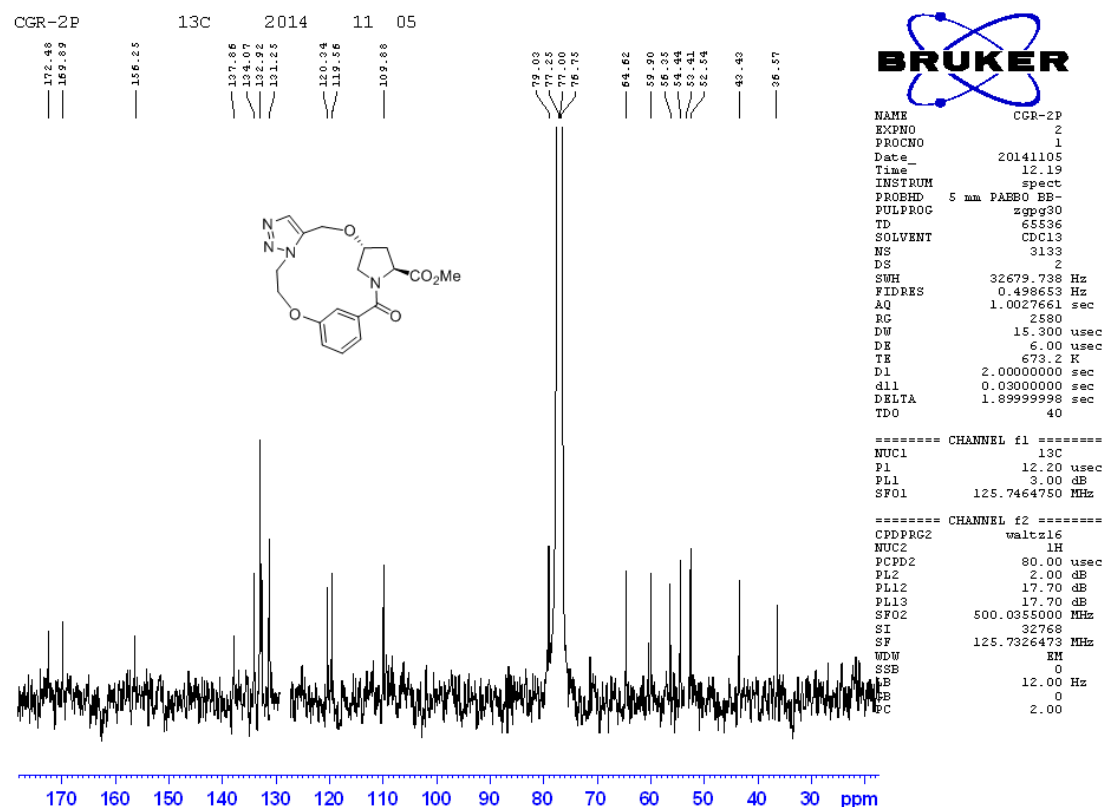


- 10c** R = OCH₃
42 R = OH
43 R = NH₂
44 R = NHPh
45 R = NHCH₂CH(CH₃)₂

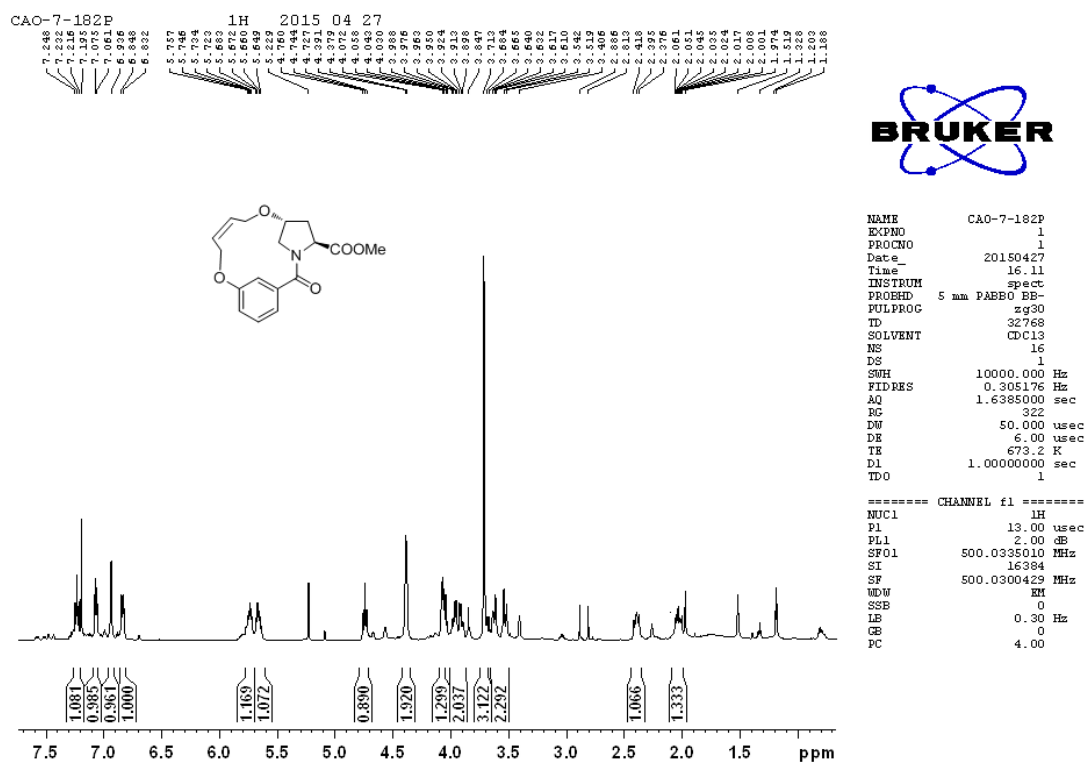
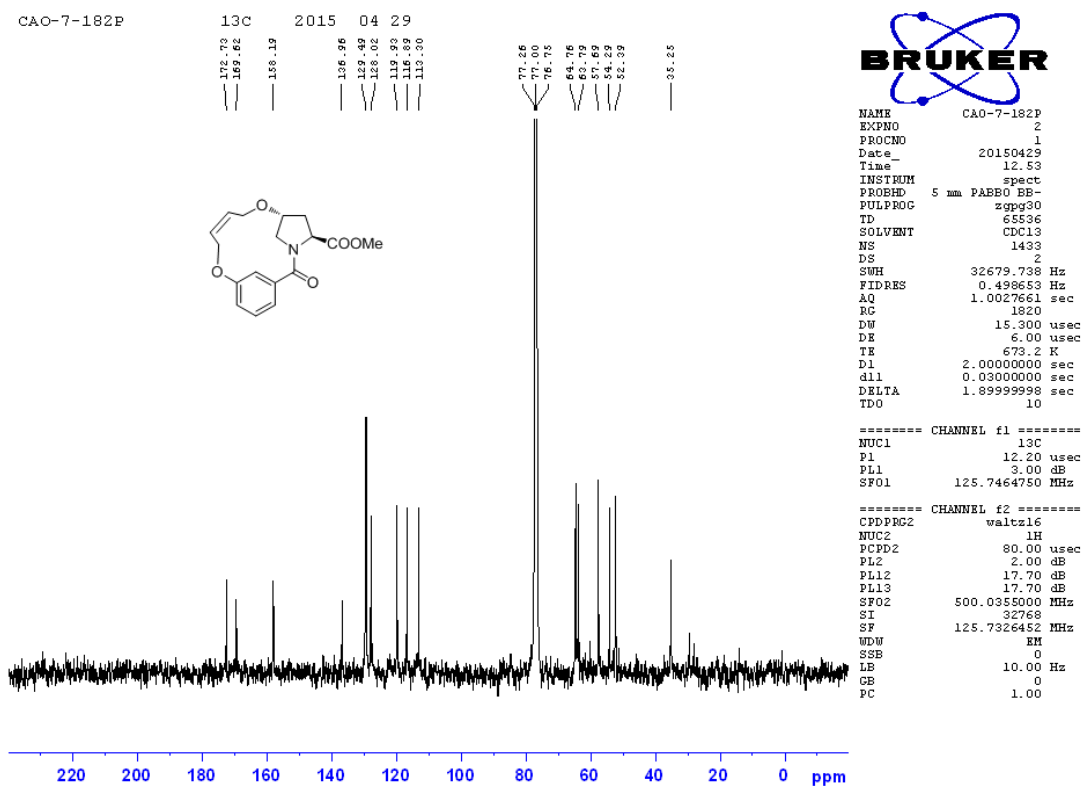
Figure S1. Target macrocycles.

Spectra of Macrocycle 6a–b

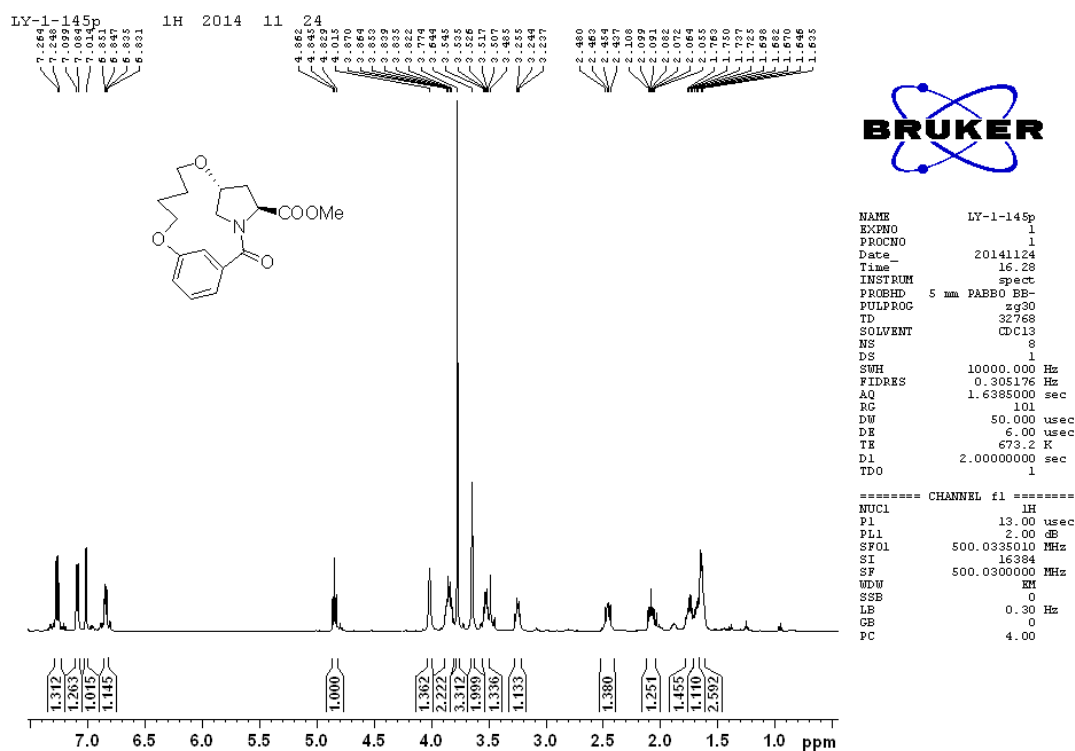
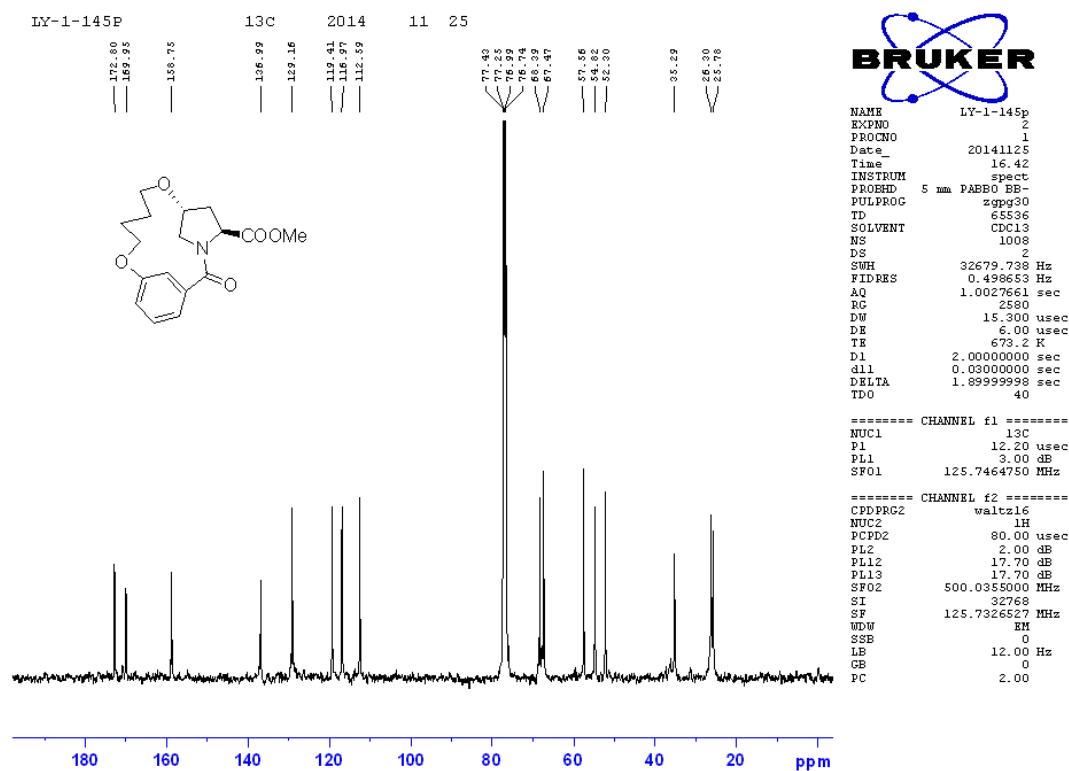
Figure S2. ¹H-NMR Spectrum of 6a.Figure S3. ¹³C-NMR Spectrum of 6a.

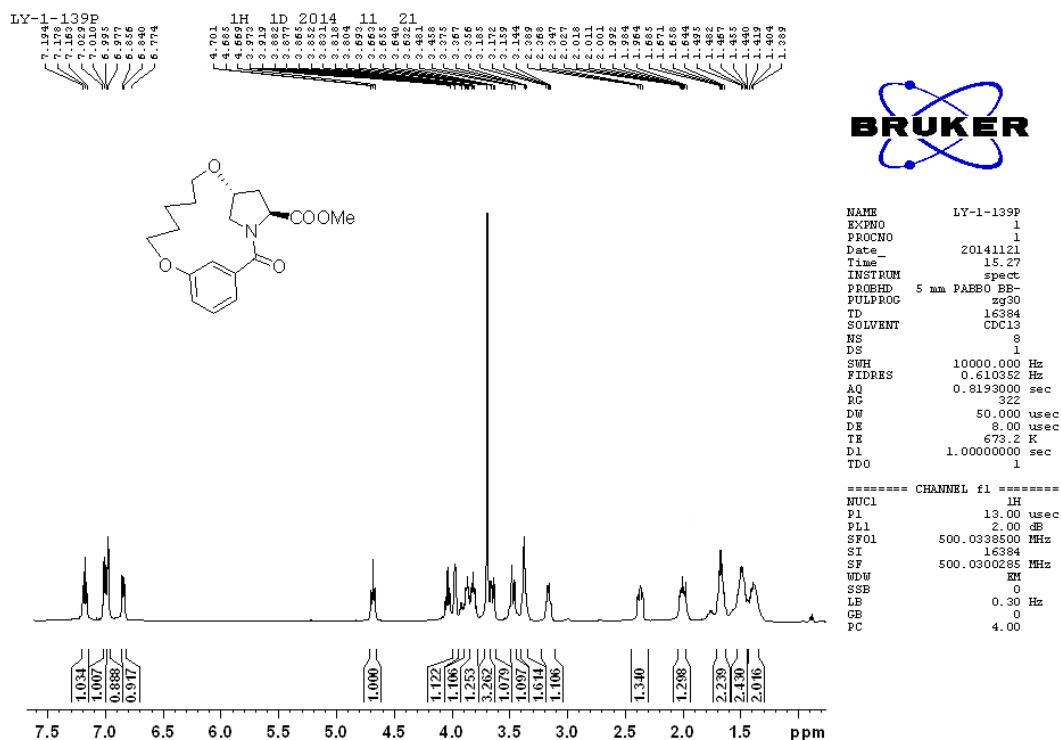
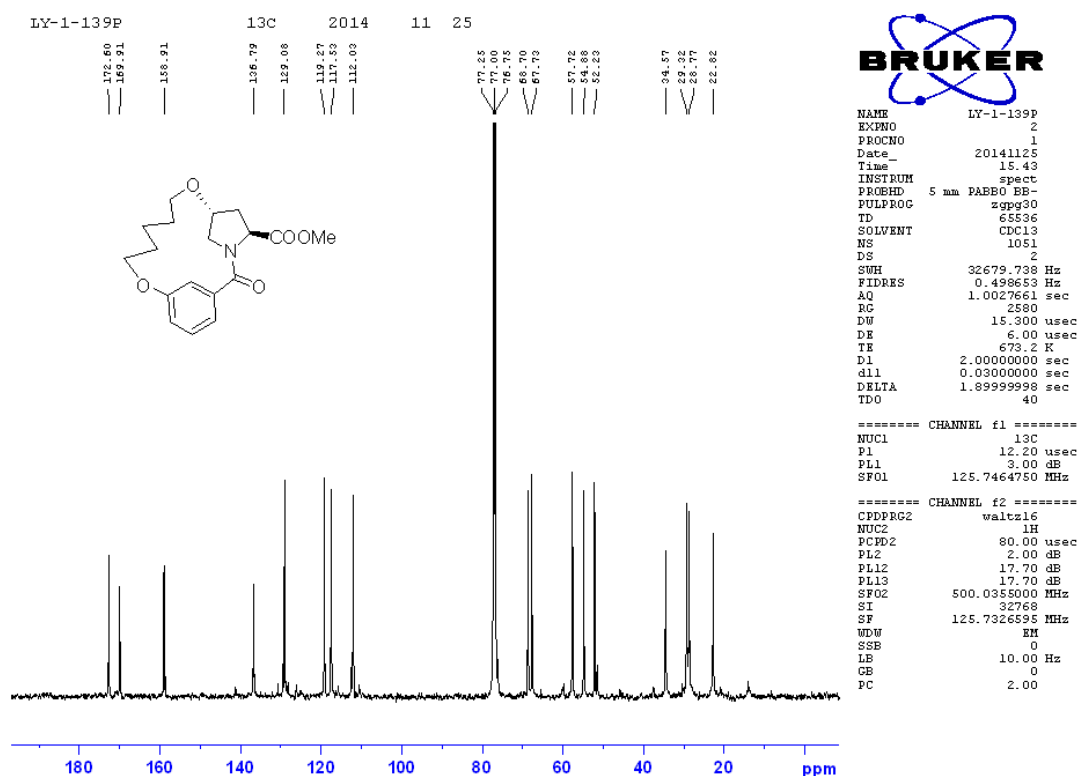
Figure S4. ¹H-NMR Spectrum of 6b.Figure S5. ¹³C-NMR Spectrum of 6b.

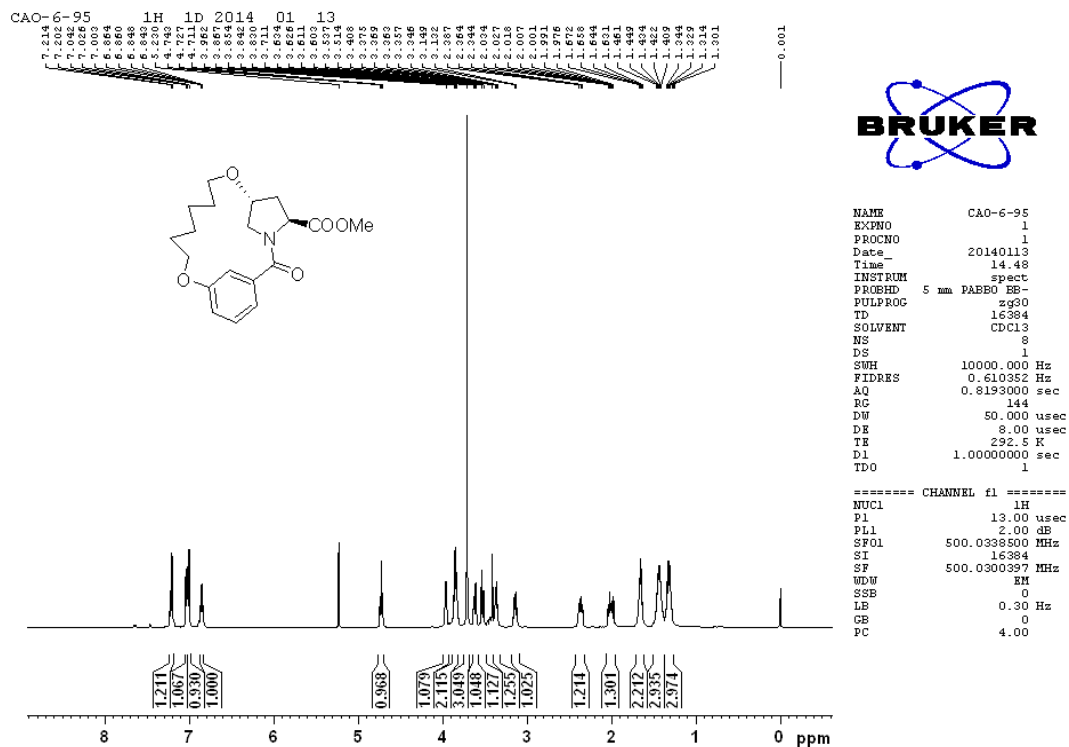
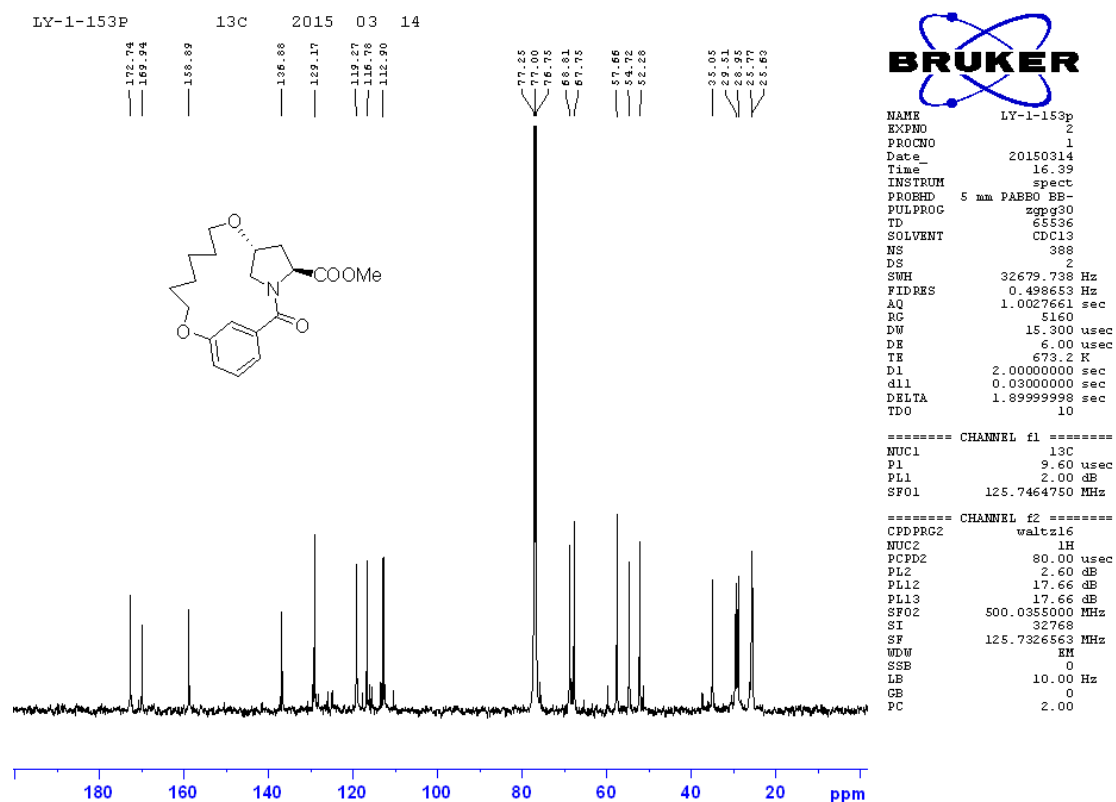
Spectra of macrocycle 9

Figure S6. ¹H-NMR Spectrum of 9.Figure S7. ¹³C-NMR Spectrum of 9.

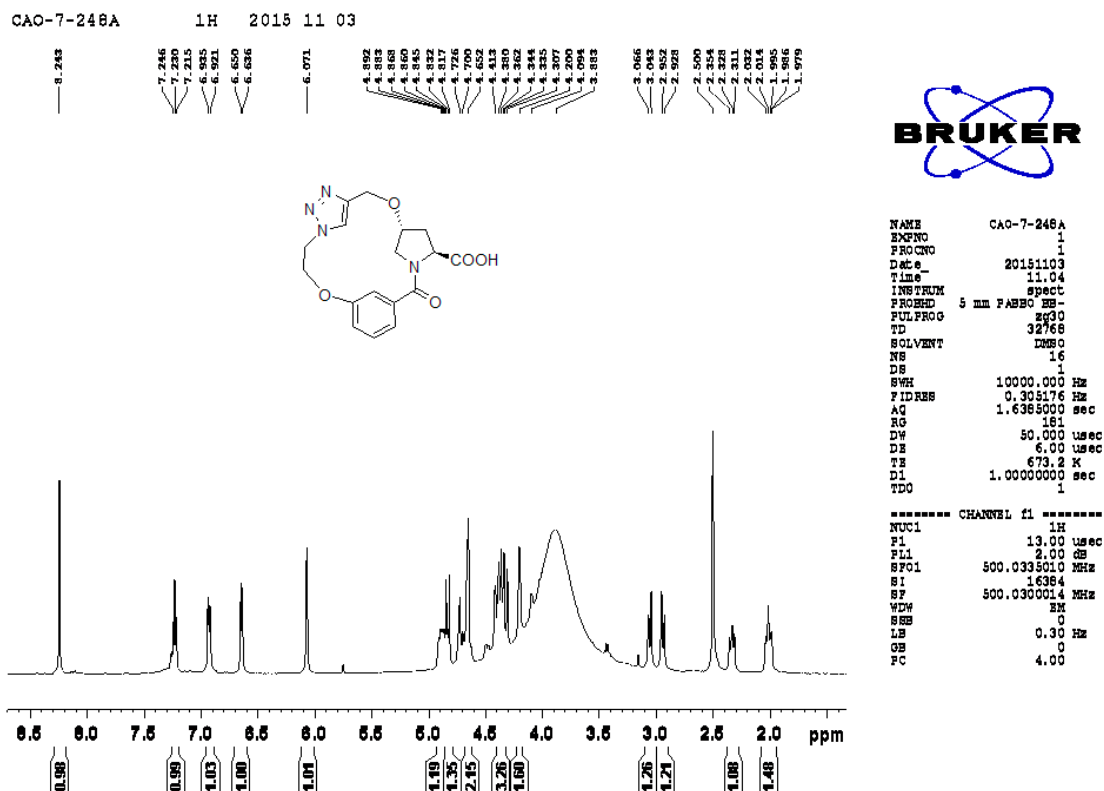
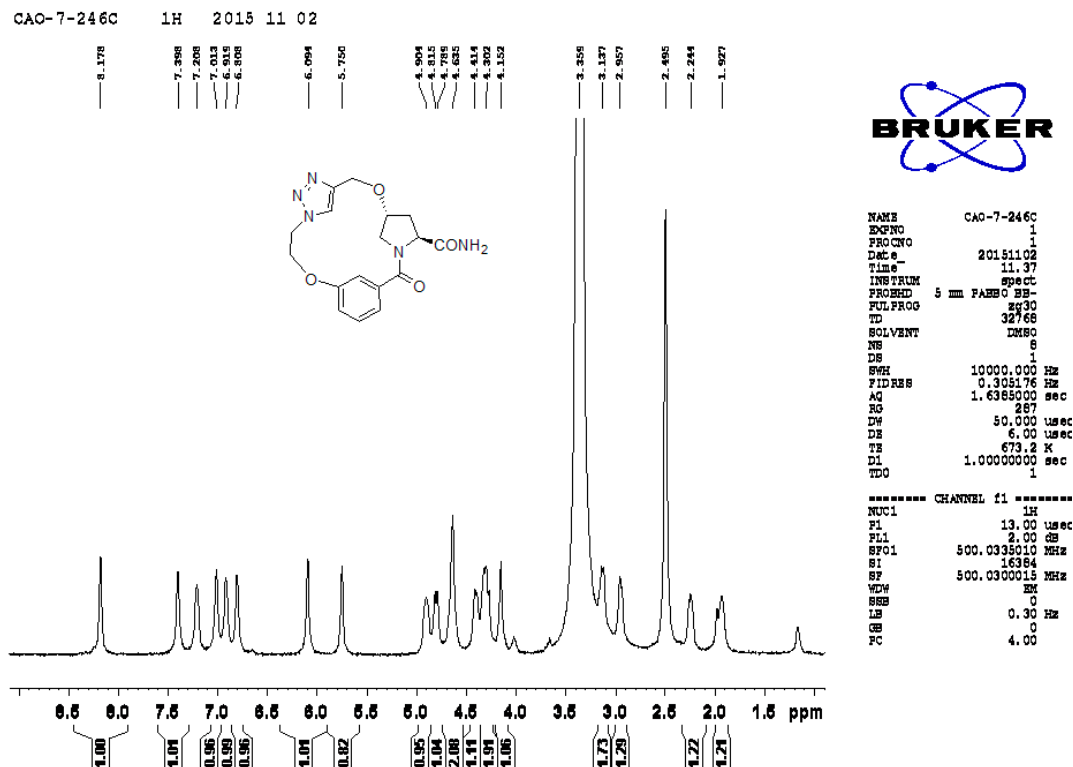
Spectra of macrocycle 10a–c

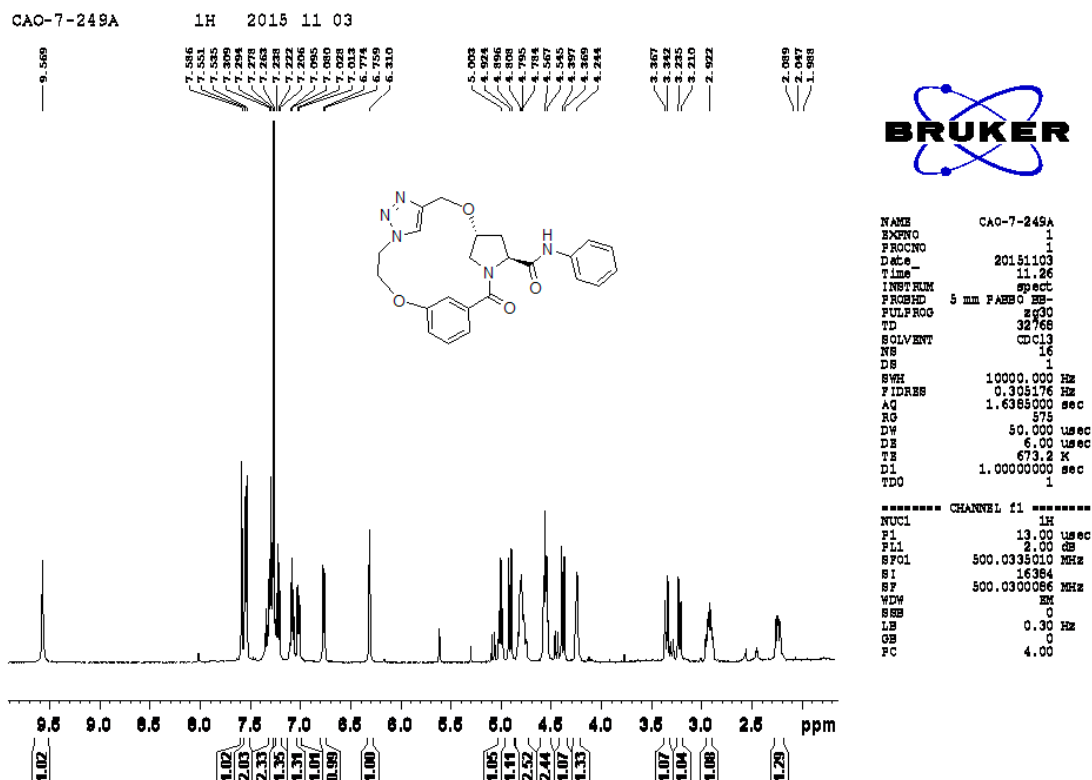
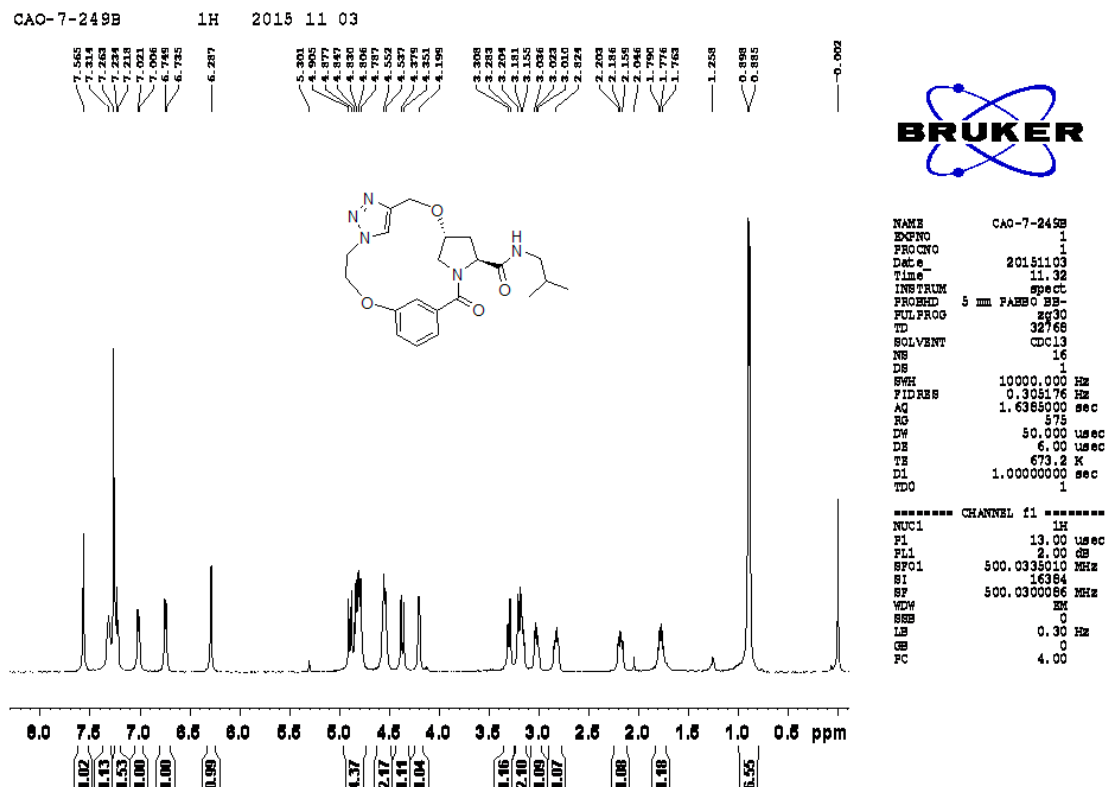
Figure S8. ¹H-NMR Spectrum of 10a.Figure S9. ¹³C-NMR Spectrum of 10a.

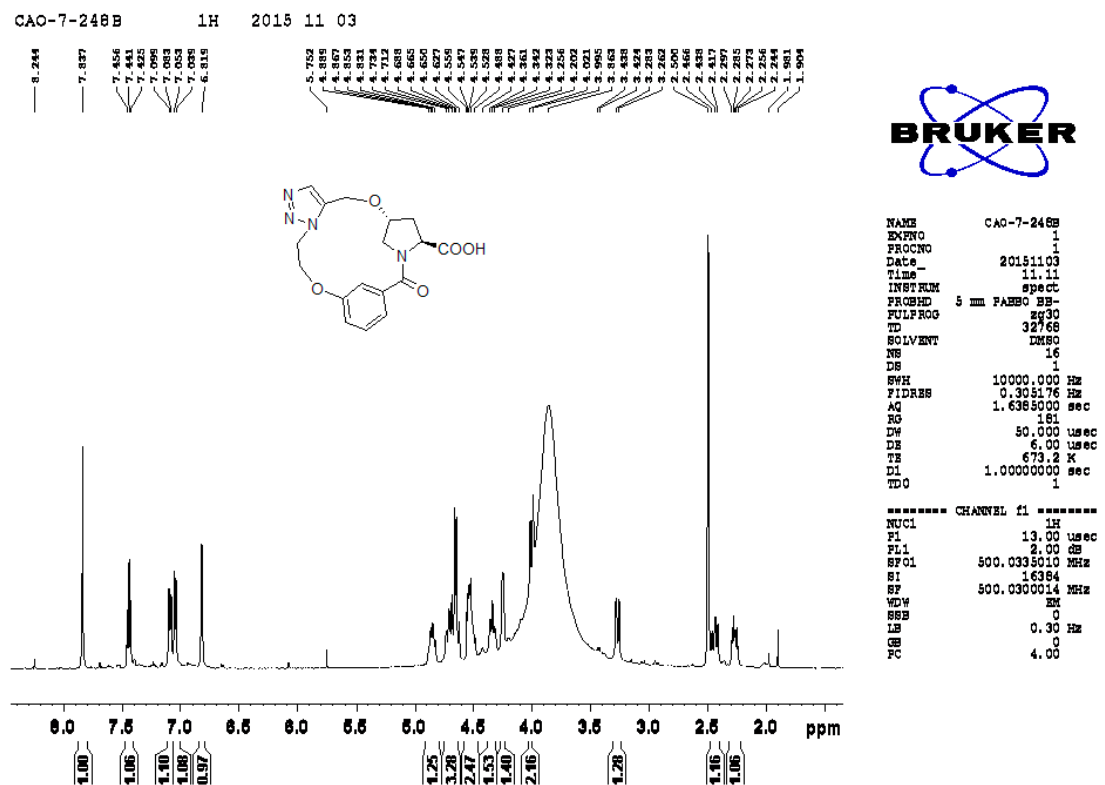
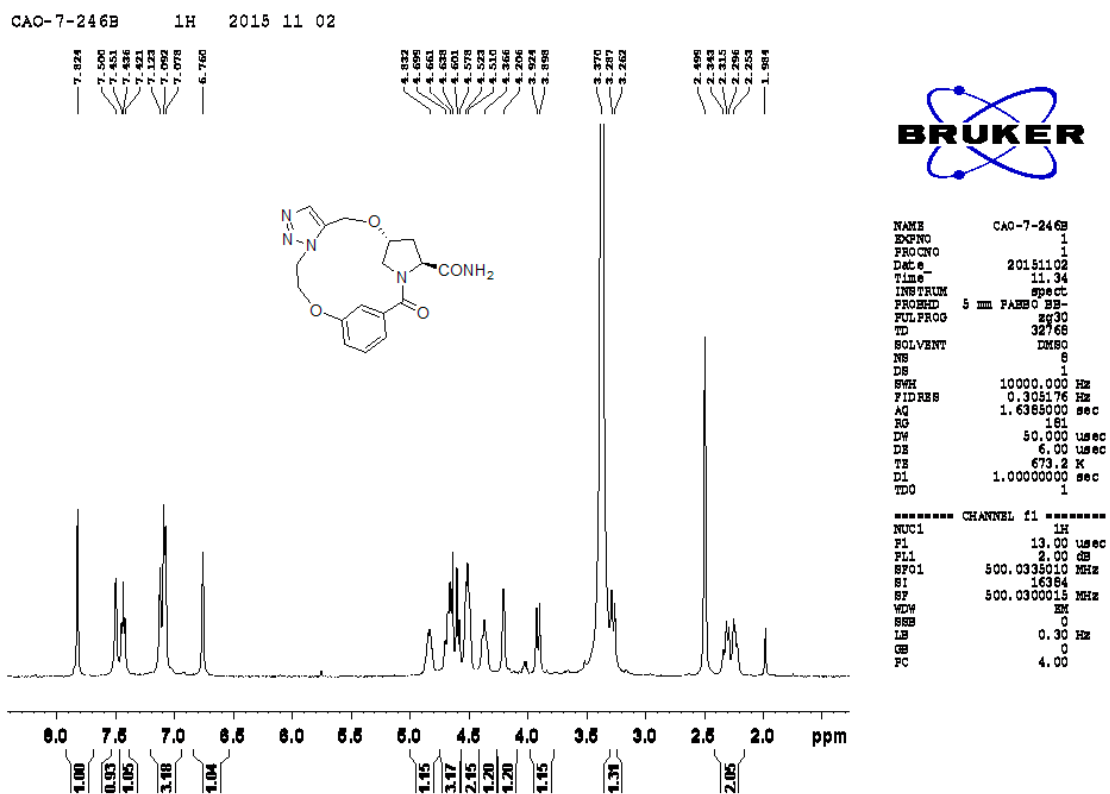
Figure S10. ¹H-NMR Spectrum of 10b.Figure S11. ¹³C-NMR Spectrum of 10b.

Figure S12. ¹H-NMR Spectrum of 10c.Figure S13. ¹³C-NMR Spectrum of 10c.

Spectra of macrocycle 22–45

Figure S14. ¹H-NMR Spectrum of 22.Figure S15. ¹H-NMR Spectrum of 23.

Figure S16. ¹H-NMR Spectrum of 24.Figure S17. ¹H-NMR Spectrum of 25.

Figure S18. ¹H-NMR Spectrum of 26.Figure S19. ¹H-NMR Spectrum of 27.

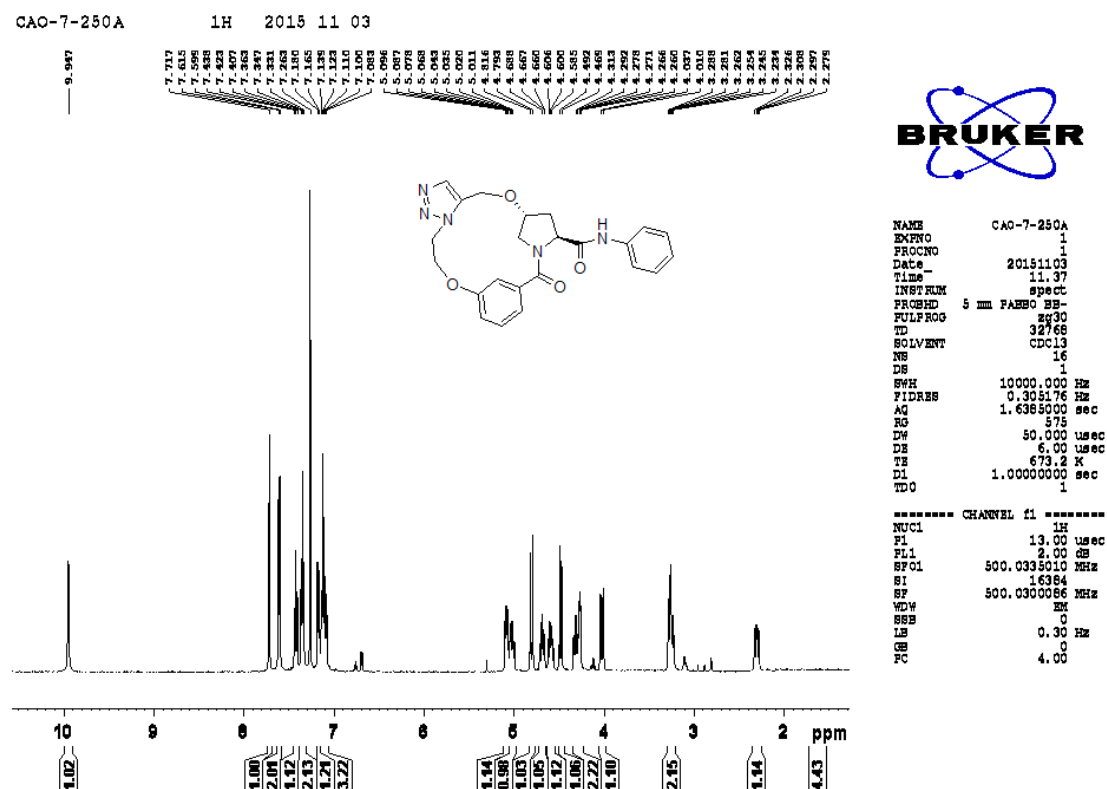


Figure S20. ^1H -NMR Spectrum of 28.

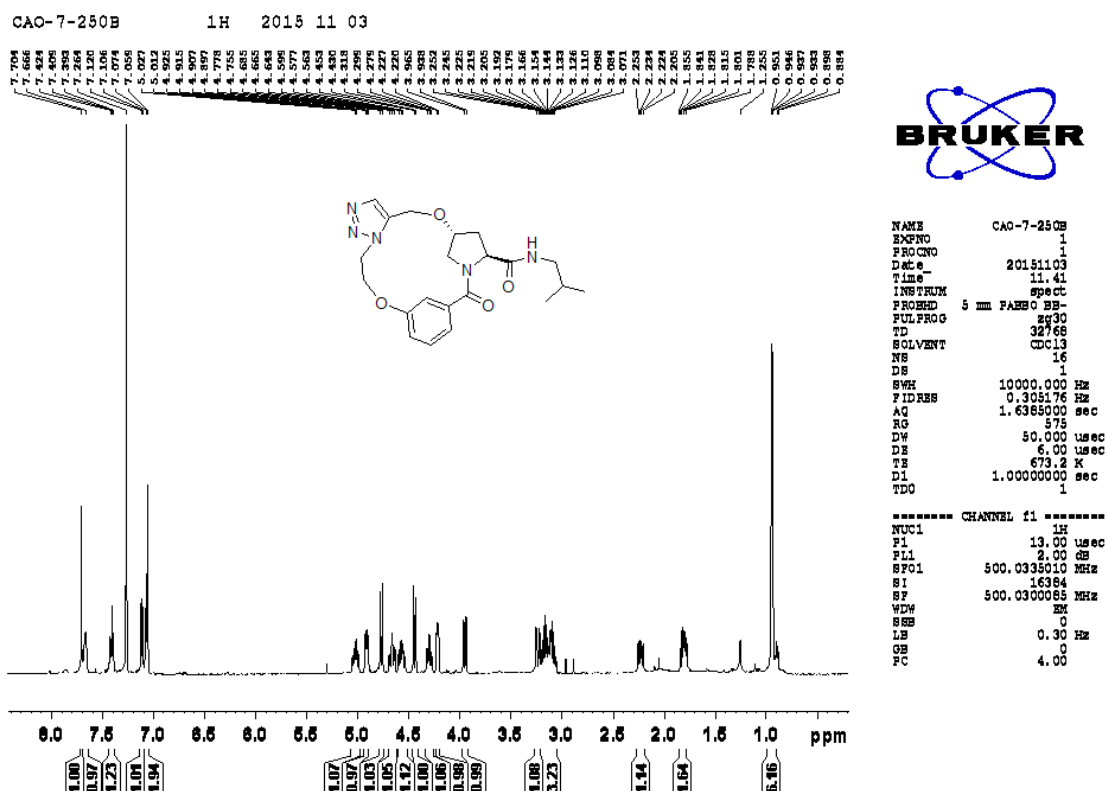
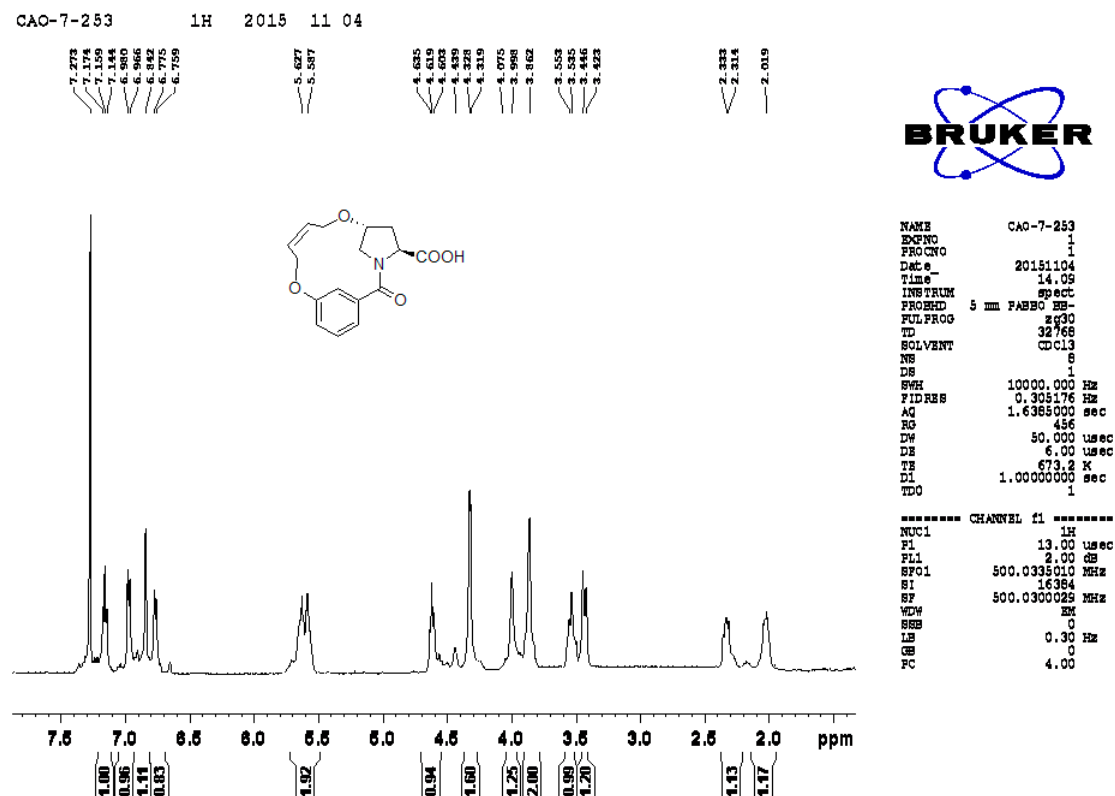
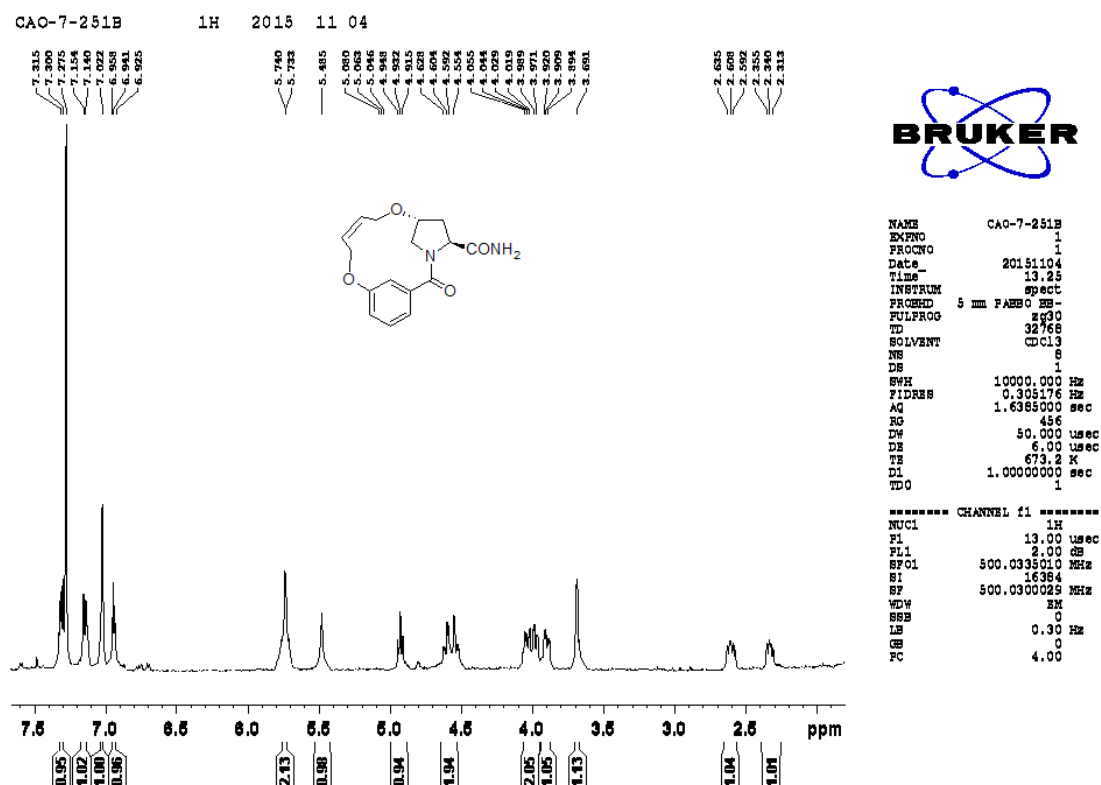
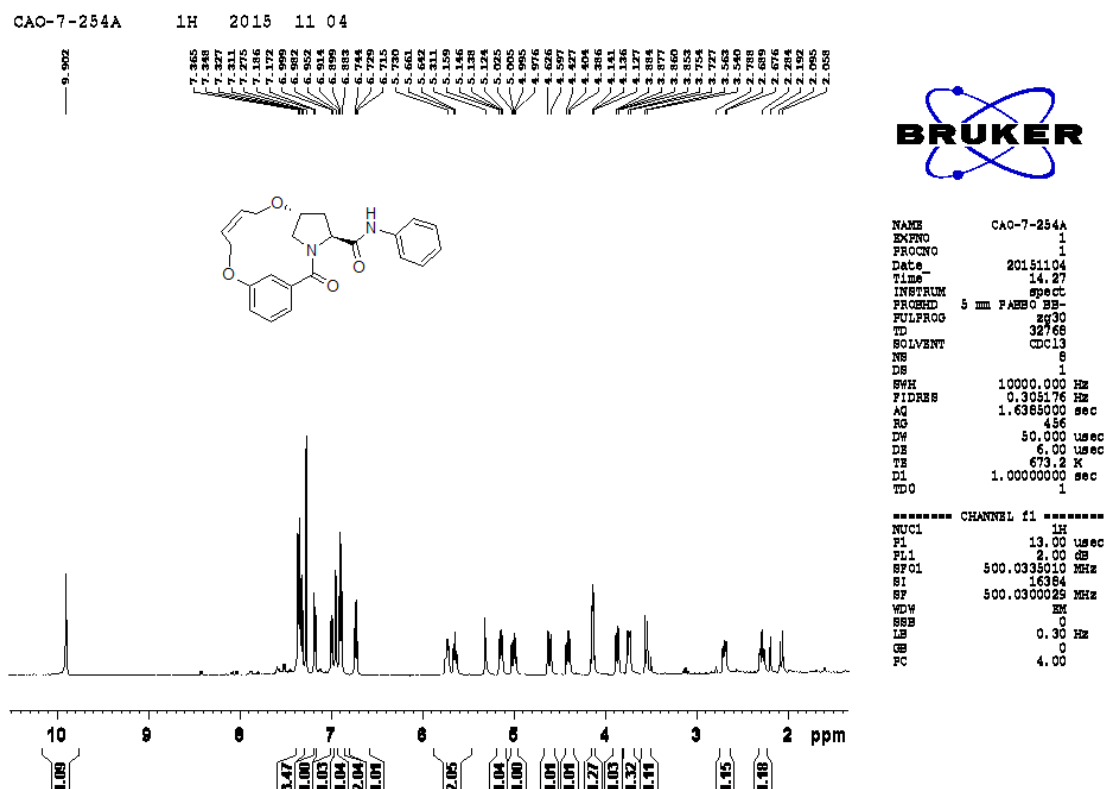
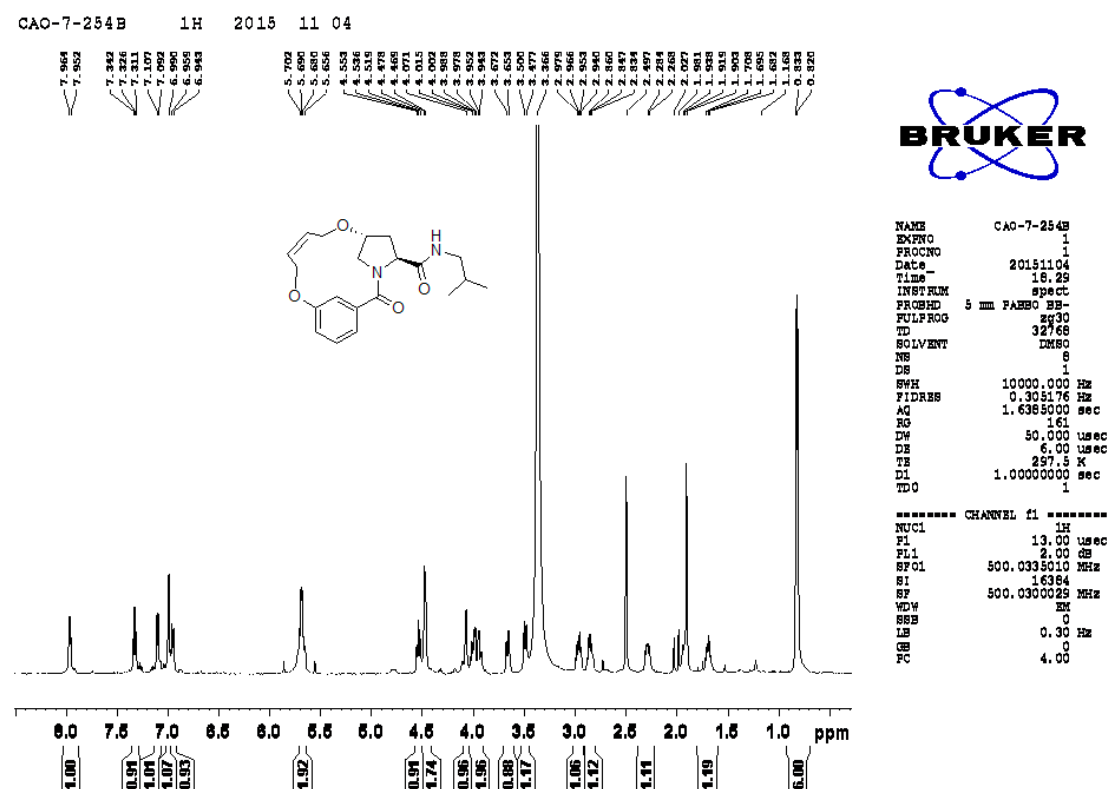
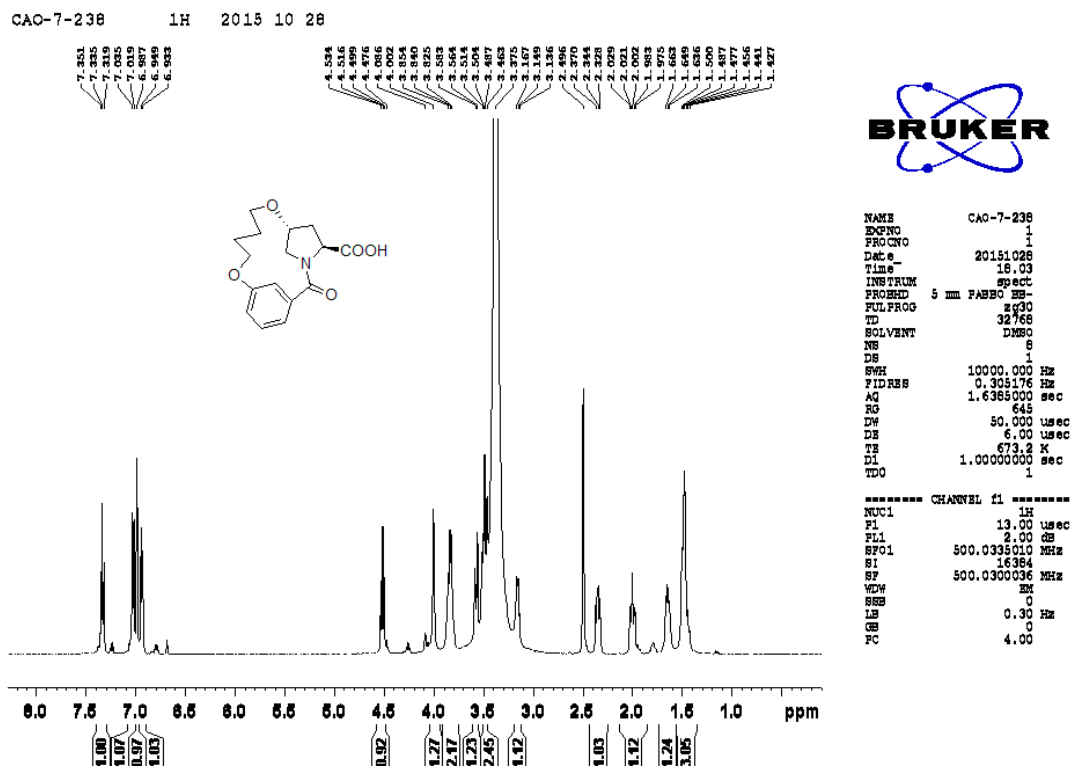
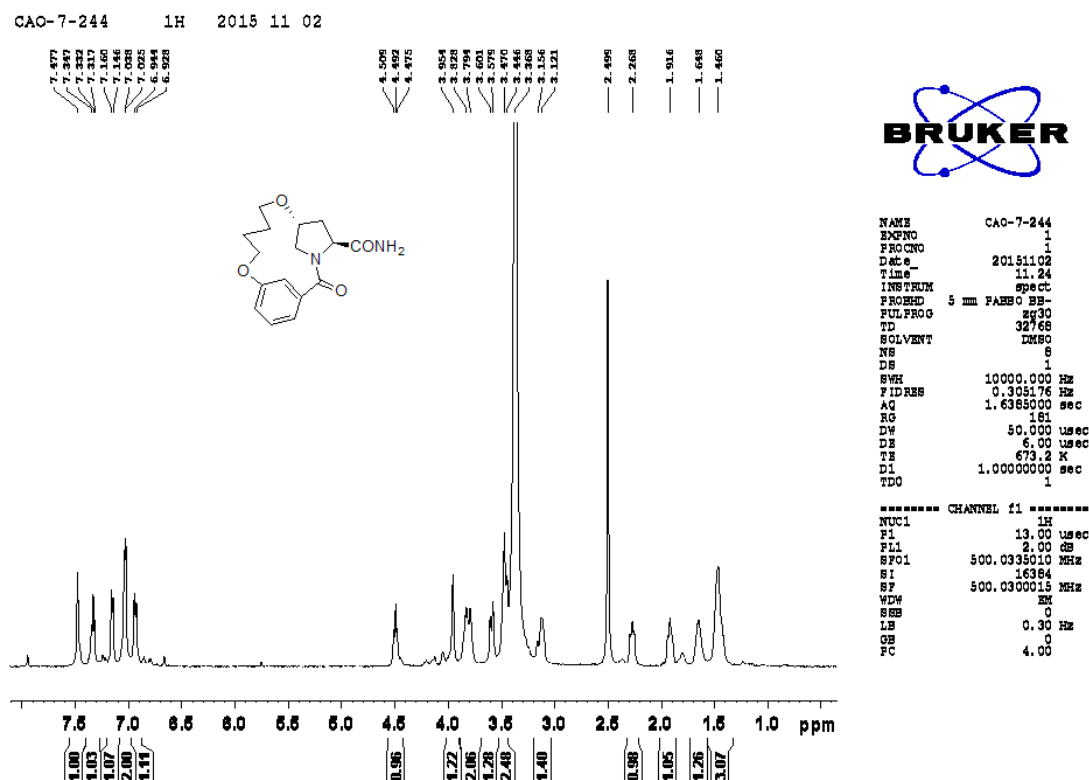
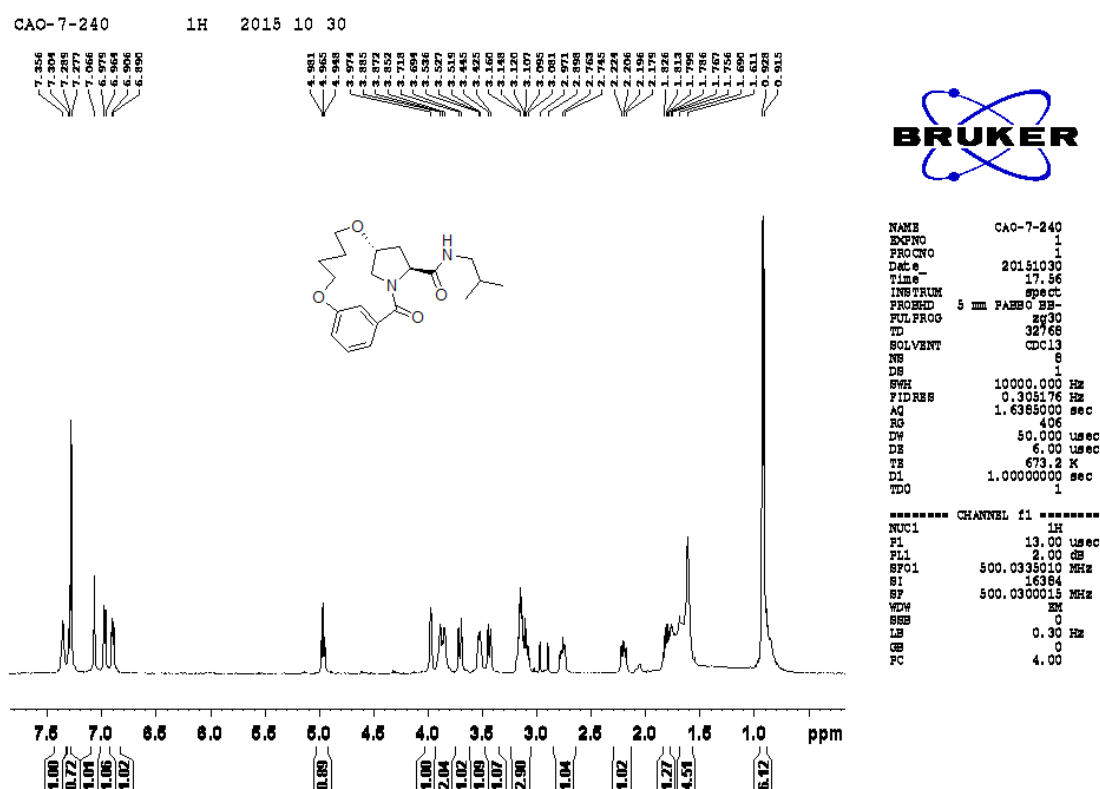
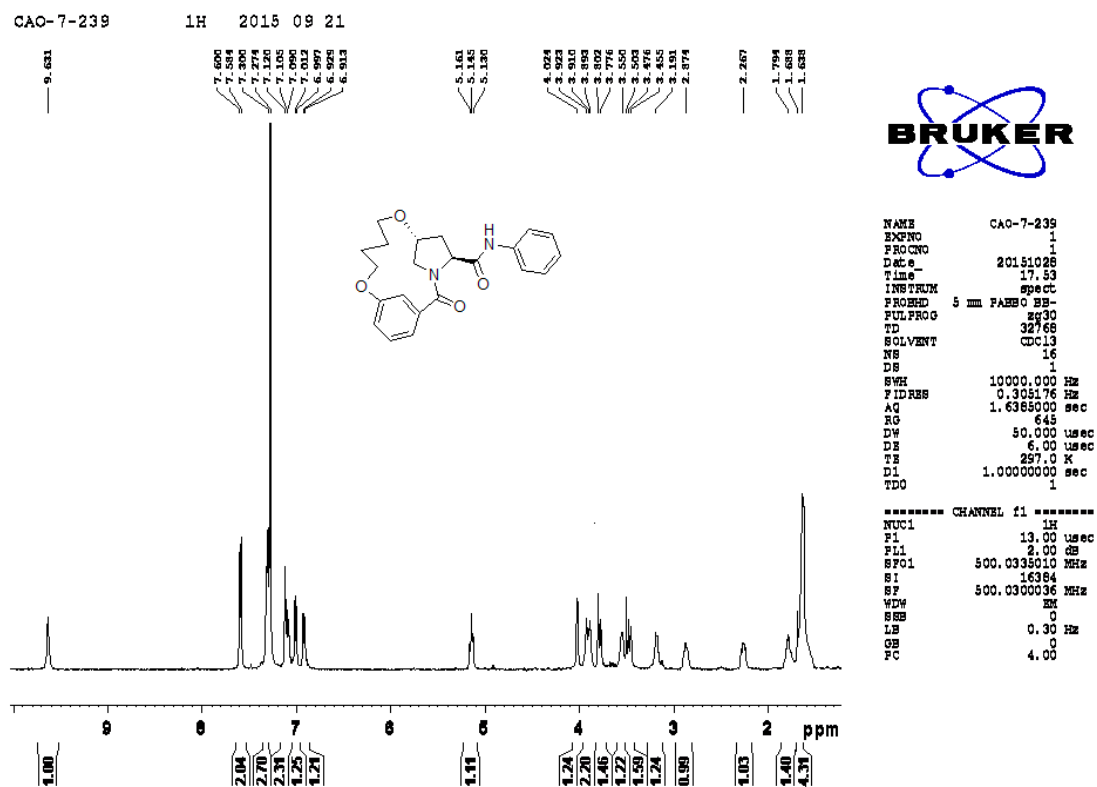


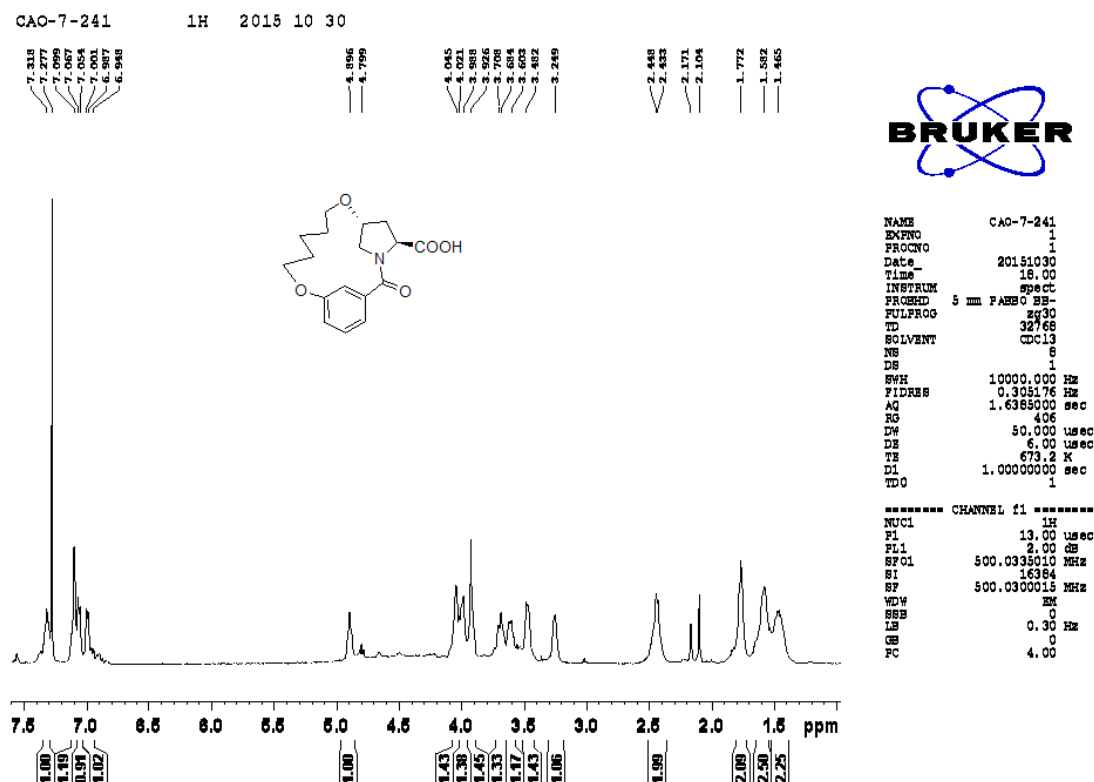
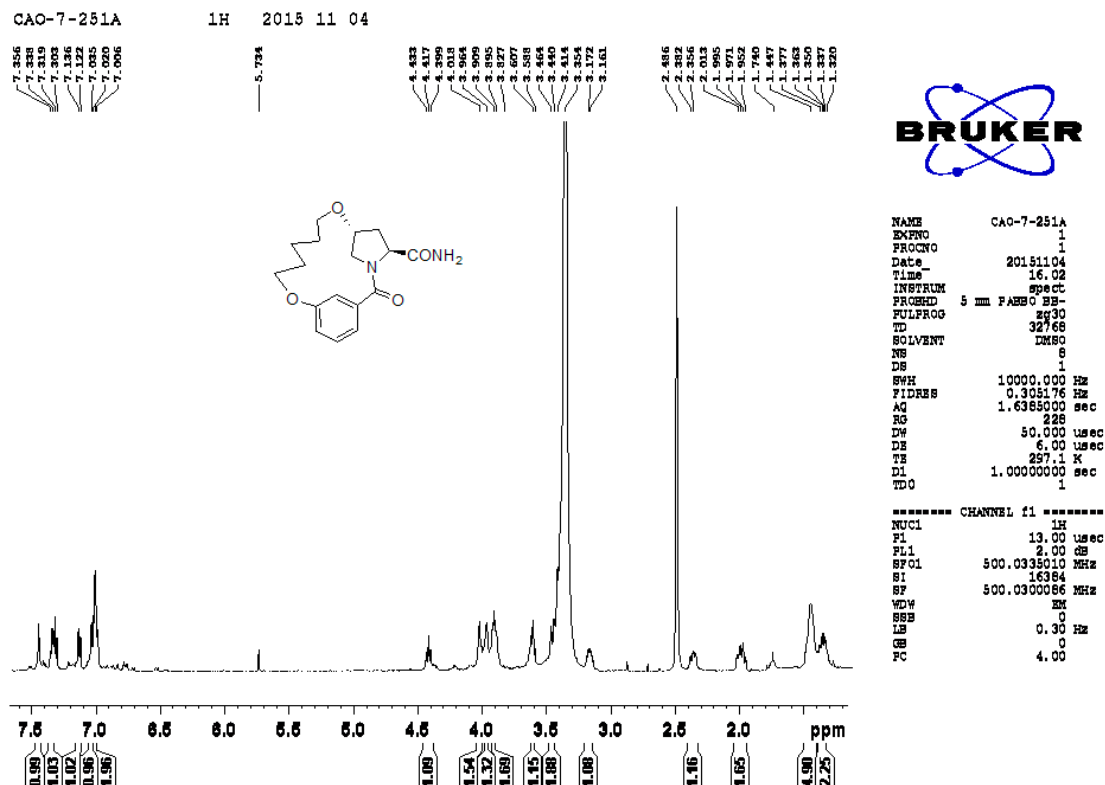
Figure S21. ^1H -NMR Spectrum of 29.

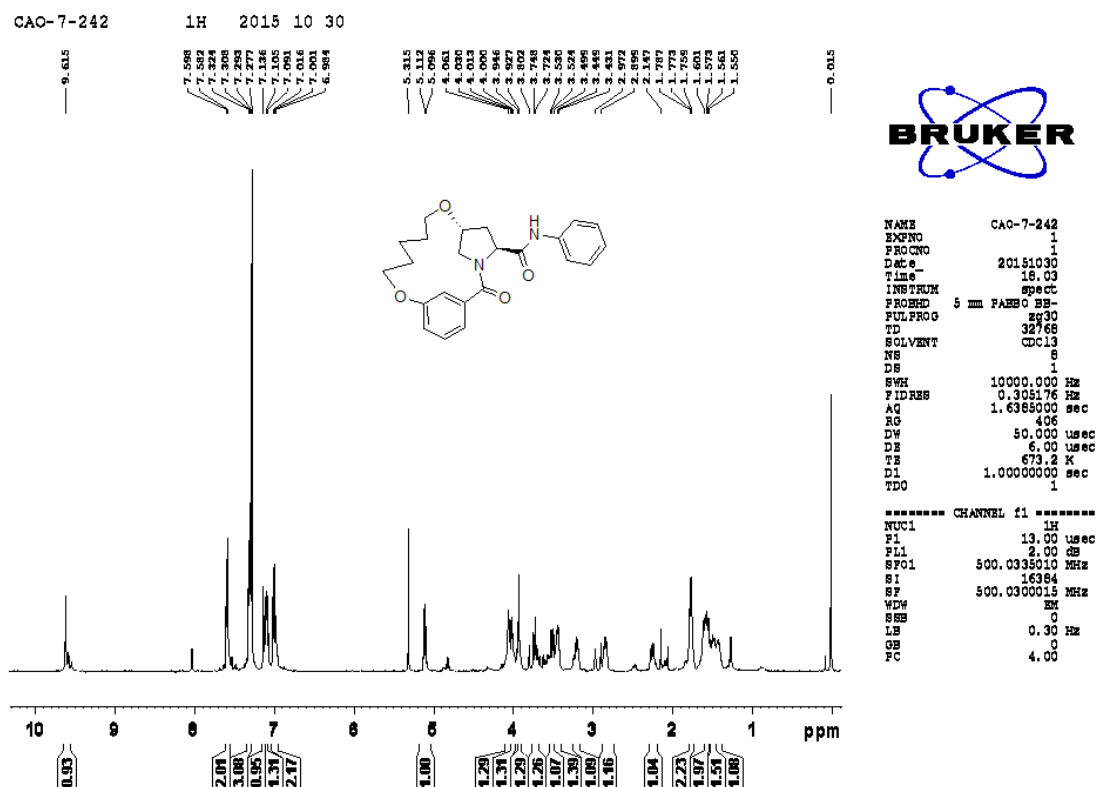
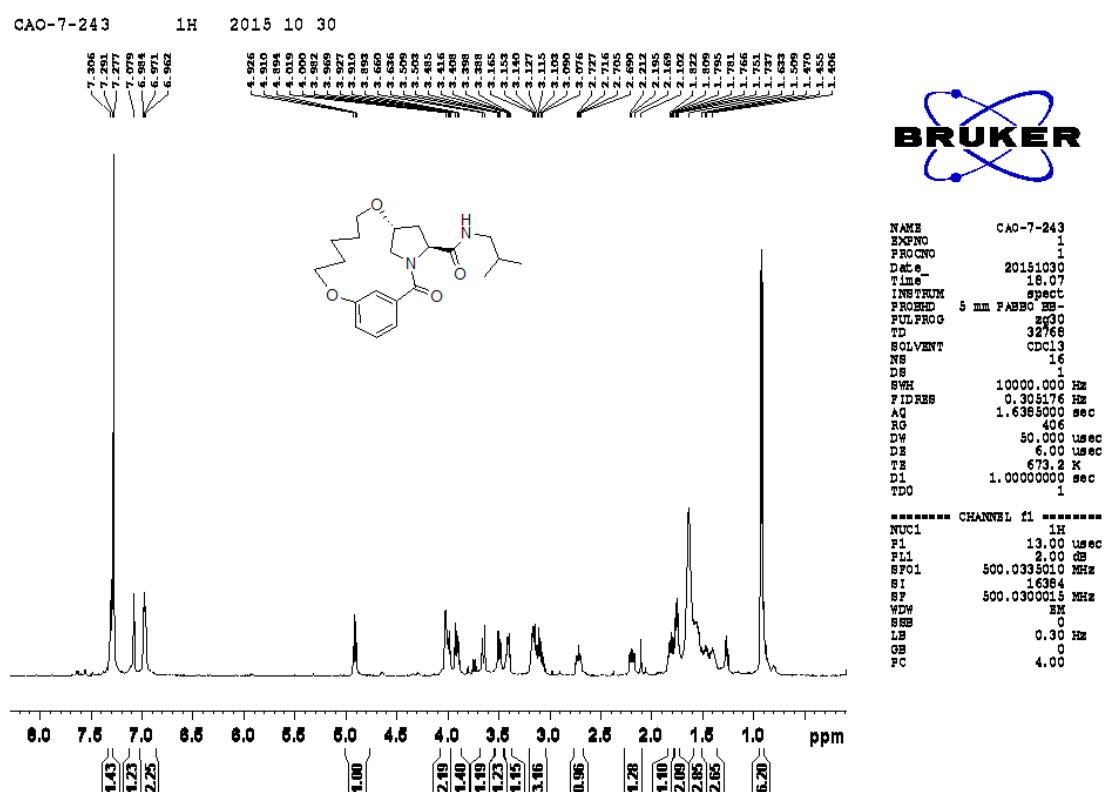
Figure S22. ¹H-NMR Spectrum of 30.Figure S23. ¹H-NMR Spectrum of 31.

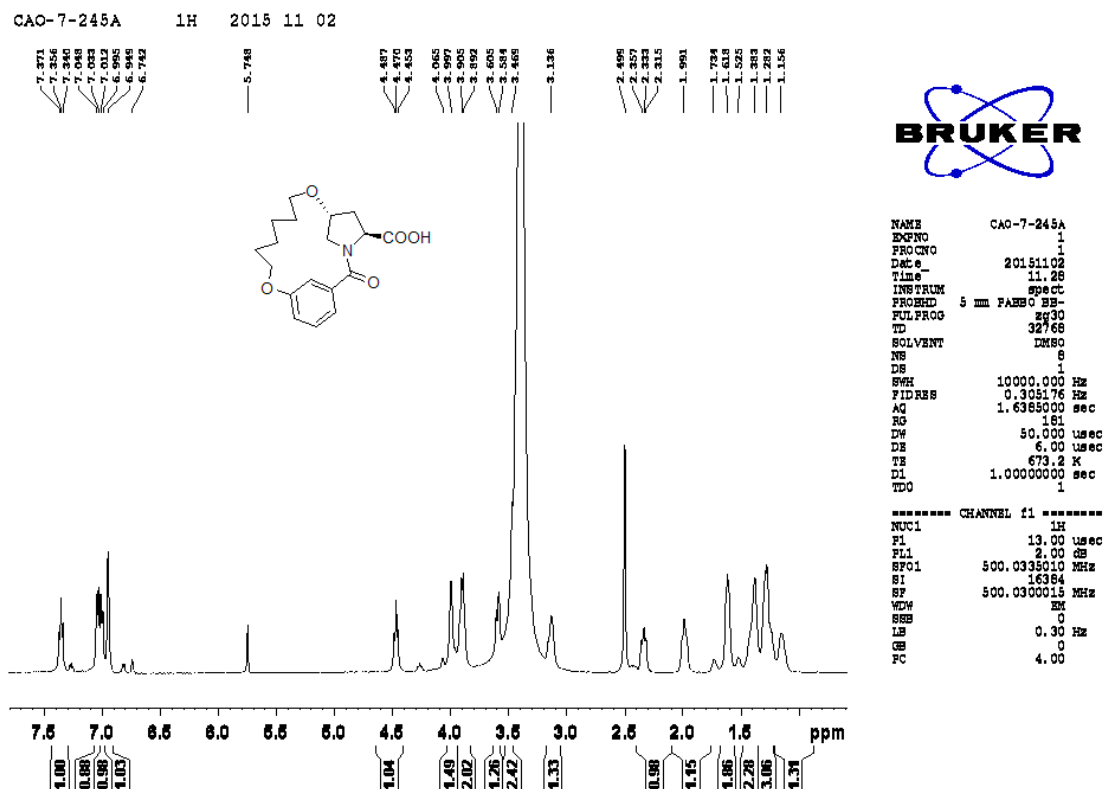
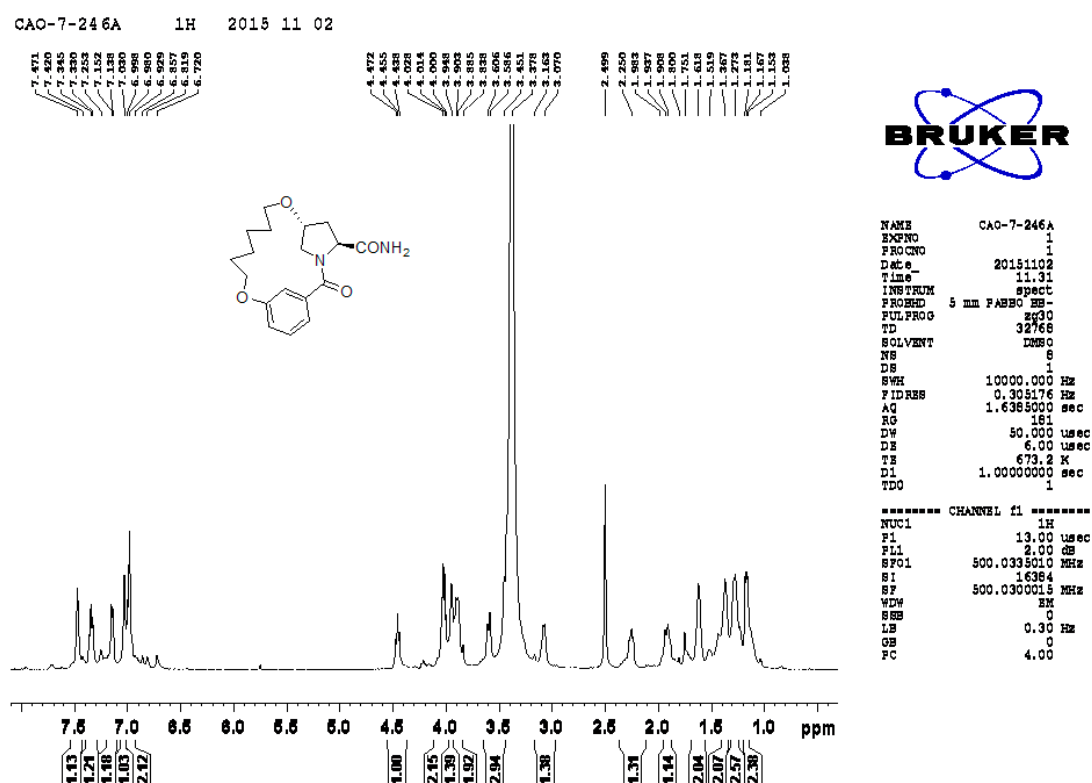
Figure S24. ¹H-NMR Spectrum of 32.Figure S25. ¹H-NMR Spectrum of 33.

Figure S26. ¹H-NMR Spectrum of 34.Figure S27. ¹H-NMR Spectrum of 35.



Figure S30. ¹H-NMR Spectrum of 38.Figure S31. ¹H-NMR Spectrum of 39.

Figure S32. ¹H-NMR Spectrum of 40.Figure S33. ¹H-NMR Spectrum of 41.

Figure S34. ¹H-NMR Spectrum of 42.Figure S35. ¹H-NMR Spectrum of 43.

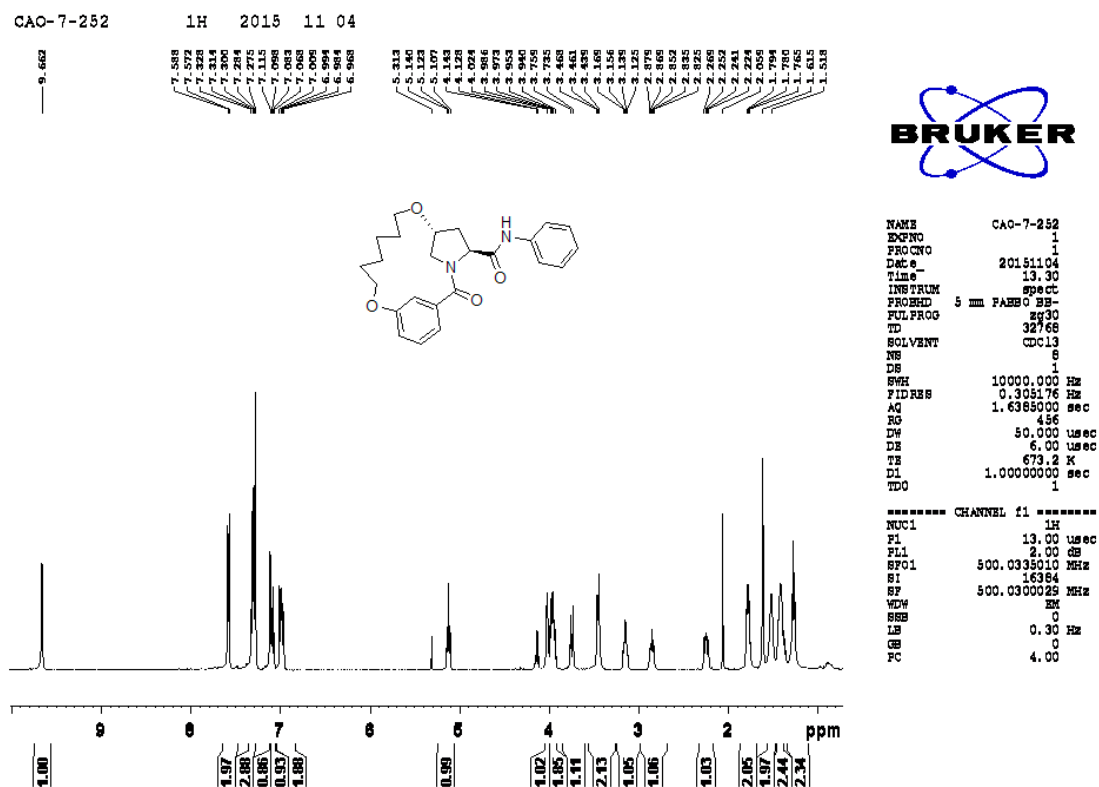


Figure S36. ^1H -NMR Spectrum of **44**.

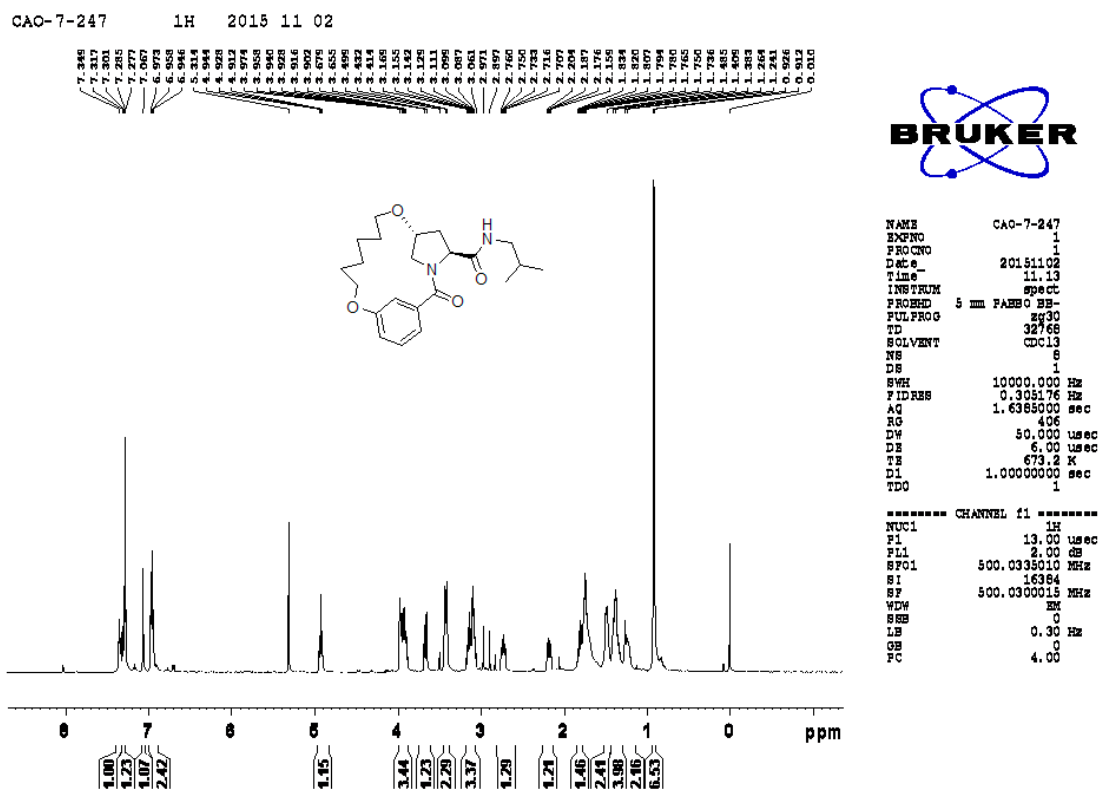


Figure S37. ^1H -NMR Spectrum of 45.