

Supplementary Information

Direct synthesis of tetrahydropyran-4-ones via O₃ReOH-catalyzed Prins cyclization of 3-chlorohomoallylic alcohols

Kwanruthai Tadpetch ^{a*}, Pongsit Vijitphan ^{a,b}, Sasiwan Kaewsen ^a, Aticha Thiraporn ^a,
Vatcharin Rukachaisirikul ^{a*}

^a *Division of Physical Science and Center of Excellence for Innovation in Chemistry, Faculty of Science, Prince of Songkla University, Hat Yai, Songkhla 90110, Thailand*

^b *Current address: Department of Chemistry, Laboratory of Organic Synthesis, Chulabhorn Research Institute, Bangkok 10210, Thailand*

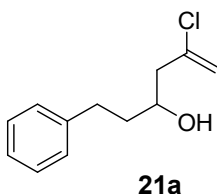
* Corresponding author. Tel.: +66 74 288437; fax: +66 74 558403.
Email address: kwanruthai.t@psu.ac.th (K. Tadpetch).

List of supplementary information

	Page
1. General Experimental Details	2
2. Characterization Data	2
3. ¹ H and ¹³ C NMR Spectra	15
4. NOE Spectra of Selected Compounds	45
5. References	46

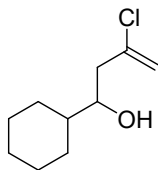
Unless otherwise stated, all reactions were conducted under a nitrogen or argon atmosphere in oven- or flamed-dried glassware. Solvents were used as received from suppliers or distilled prior to use using standard procedures. Perrhenic acid solution (75-80 wt. % in H₂O) was purchased from Sigma-Aldrich. All other reagents were obtained from commercial sources and used without further purification. Chlorohomoallylic alcohols **21a**, **21b** and **21c** were prepared following Li's protocol¹ using 2-chloro-2-propene and dihydrocinnamaldehyde, cyclohexanecarboxaldehyde and benzaldehyde respectively. Column chromatography was performed on Silica gel 60 (0.063-0.200 mm, Merck). Thin-layer chromatography (TLC) was performed on Silica gel 60 F₂₅₄ (Merck). ¹H and ¹³C NMR spectroscopic data were recorded at ambient temperature on 300 or 500 MHz Bruker FTNMR Ultra Shield spectrometers. ¹H NMR spectra are reported in ppm on the δ scale and referenced to the internal tetramethylsilane. The data are presented as follows: chemical shift, multiplicity (s = singlet, d = doublet, t = triplet, q = quartet, m = multiplet, br = broad, app = apparent), coupling constant(s) in hertz (Hz), and integration. ¹³C NMR spectra were reported in ppm relative to CDCl₃ (77.16 ppm). Infrared (IR) spectra were recorded on a Perkin Elmer 783 FTS165 FT-IR spectrometer. High-resolution mass spectra were obtained on a Ultra-Performance Liquid Chromatography–High Resolution Mass Spectrometer (Agilent LC-QTOF 6500 system), Mae Fah Luang University or a High-Performance Liquid Chromatograph–Mass Spectrometer (Shimadzu LCMS-IT-TOF Model LC-20ADXR), Thammasat University. Melting points were measured using an Electrothermal IA9200 melting point apparatus and are uncorrected.

2. Characterization Data



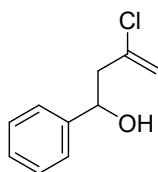
5-Chloro-1-phenylhex-5-en-3-ol (21a). Light yellow oil: $R_f = 0.50$ (20% EtOAc/hexanes); ¹H NMR (300 MHz, CDCl₃) δ 7.38–7.24 (m, 2H), 7.24–7.12 (m, 3H), 5.28 (d, $J = 0.9$ Hz, 1H), 5.24 (app s, 1H), 3.97 (tt, $J = 6.6, 6.0$ Hz, 1H), 2.90–2.60 (m, 2H), 1.92–1.76 (m, 2H); ¹³C NMR (75 MHz, CDCl₃) δ 141.8, 139.3, 128.47, 128.45,

126.0, 115.1, 68.2, 47.3, 38.2, 32.0. ^1H and ^{13}C NMR data matched those previously reported by Leighton et al.²



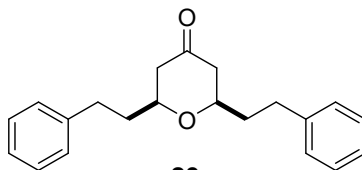
21b

3-Chloro-1-cyclohexylbut-3-en-1-ol (21b). Light yellow oil: $R_f = 0.55$ (20% EtOAc/hexanes); ^1H NMR (300 MHz, CDCl_3) δ 5.28 (s, 1H), 5.26 (s, 1H), 3.71 (ddd, $J = 9.0, 5.4, 3.6$ Hz, 1H), 2.51 (ddd, $J = 14.4, 3.3, 0.9$ Hz, 1H), 2.41 (dd, $J = 14.4, 9.0$ Hz, 1H), 1.92–1.66 (m, 5H), 1.45–1.33 (m, 1H), 1.32–0.98 (m, 4H); ^{13}C NMR (75 MHz, CDCl_3) δ 140.2, 115.0, 72.5, 44.4, 43.1, 29.2, 27.9, 26.5, 26.3, 26.2; IR (thin film): 3384, 2926, 2853, 1636, 1449, 881 cm^{-1} .



21c

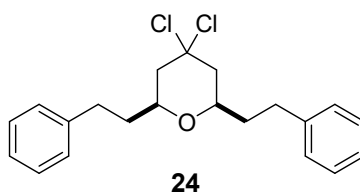
3-Chloro-1-phenylbut-3-en-1-ol (21c). Light yellow oil: $R_f = 0.65$ (20% EtOAc/hexanes); ^1H NMR (300 MHz, CDCl_3) δ 7.36–7.22 (m, 5H), 5.24 (d, $J = 1.2$ Hz, 1H), 5.18 (app s, 1H), 4.98 (dd, $J = 8.7, 4.5$ Hz, 1H), 2.73 (dd, $J = 14.4, 8.7$ Hz, 1H), 2.61 (ddd, $J = 14.4, 4.5, 0.6$ Hz, 1H), 2.38 (br s, 1H); ^{13}C NMR (75 MHz, CDCl_3) δ 143.1, 138.9, 128.6, 127.9, 125.9, 115.5, 71.2, 49.2. ^1H and ^{13}C NMR data matched those previously reported by Antilla et al.³



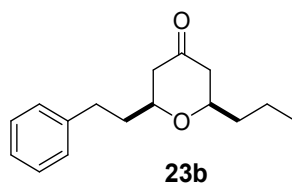
23a

2,6-Diphenethyltetrahydropyran-4-one (23a). Light yellow oil: $R_f = 0.42$ (20% EtOAc/hexanes); ^1H NMR (300 MHz, CDCl_3) δ 7.36–7.27 (m, 4H), 7.27–7.18 (m, 6H), 3.62–3.49 (m, 2H), 2.99–2.85 (m, 2H), 2.85–2.71 (m, 2H), 2.42–2.22 (m, 2H), 2.13–

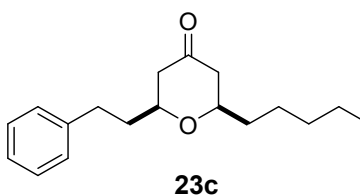
1.95 (m, 2H), 1.92–1.70 (m, 2H); ^{13}C NMR (75 MHz, CDCl_3) δ 207.3, 141.5, 128.6, 128.5, 126.1, 75.9, 48.0, 38.1, 31.7. ^1H and ^{13}C NMR data matched those previously reported by Scheidt et al.⁴



4,4-Dichloro-2,6-diphenethyltetrahydropyran (24). Light yellow oil: $R_f = 0.65$ (20% EtOAc/hexanes); ^1H NMR (300 MHz, CDCl_3) δ 7.33–7.25 (m, 4H), 7.25–7.16 (m, 6H), 3.70–3.60 (m, 2H), 2.92–2.79 (m, 2H), 2.78–2.65 (m, 2H), 2.47 (d, $J = 13.8$ Hz, 2H), 2.06 (dd, $J = 13.8, 11.1$ Hz, 2H), 1.97–1.82 (m, 2H), 1.81–1.68 (m, 2H); ^{13}C NMR (75 MHz, CDCl_3) δ 141.8, 128.5, 126.0, 87.9, 73.5, 51.5, 37.0, 31.8; IR (thin film): 3026, 2925, 1456, 1093, 750, 698 cm^{-1} ; HRMS (ES/MeOH): m/z calcd for $\text{C}_{21}\text{H}_{25}\text{Cl}_2\text{O}$ ($\text{M} + \text{H}$)⁺ 363.1282, found 363.1184.

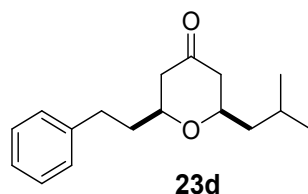


2-Phenethyl-6-propyltetrahydropyran-4-one (23b). Light yellow oil: $R_f = 0.44$ (20% EtOAc/hexanes); ^1H NMR (300 MHz, CDCl_3) δ 7.33–7.24 (m, 2H), 7.23–7.15 (m, 3H), 3.62–3.45 (m, 2H), 2.90–2.66 (m, 2H), 2.41–2.16 (m, 4H), 2.06–1.91 (m, 1H), 1.86–1.35 (m, 5H), 0.97 (d, $J = 6.9$ Hz, 3H); ^{13}C NMR (75 MHz, CDCl_3) δ 207.7, 141.5, 128.6, 128.5, 126.0, 76.8, 75.7, 48.1, 48.0, 38.7, 38.0, 31.6, 18.8, 14.0; IR (thin film): 3026, 2958, 1718, 1456, 750, 700 cm^{-1} ; HRMS (ES/MeOH): m/z calcd for $\text{C}_{16}\text{H}_{22}\text{NaO}_2$ ($\text{M} + \text{Na}$)⁺ 269.1517, found 269.1514.

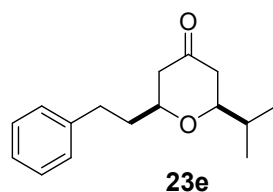


2-Pentyl-6-phenethyltetrahydropyran-4-one (23c). Light yellow oil: $R_f = 0.48$ (20% EtOAc/hexanes); ^1H NMR (300 MHz, CDCl_3) δ 7.32–7.23 (m, 2H), 7.22–7.14 (m, 3H),

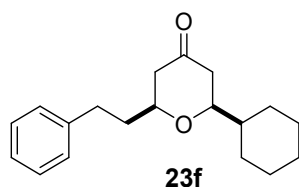
3.57–3.40 (m, 2H), 2.90–2.67 (m, 2H), 2.39–2.12 (m, 4H), 2.05–1.90 (m, 1H), 1.85–1.63 (m, 2H), 1.60–1.25 (m, 7H), 0.92 (d, $J = 6.9$ Hz, 3H); ^{13}C NMR (75 MHz, CDCl_3) δ 207.6, 141.5, 128.5, 126.0, 77.0, 75.7, 48.1, 48.0, 38.0, 36.5, 31.7, 31.6, 25.2, 22.6, 14.1; IR (thin film): 2930, 2859, 1719, 1456, 1055, 700 cm^{-1} ; HRMS (ES/MeOH): m/z calcd for $\text{C}_{18}\text{H}_{26}\text{NaO}_2$ ($\text{M} + \text{Na}$) $^+$ 297.1830, found 297.1794.



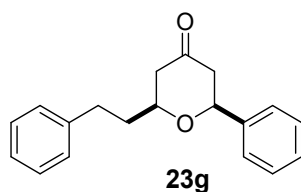
2-Isobutyl-6-phenethyltetrahydropyran-4-one (23d). Light yellow oil: $R_f = 0.46$ (20% EtOAc/hexanes); ^1H NMR (300 MHz, CDCl_3) δ 7.32–7.23 (m, 2H), 7.22–7.14 (m, 3H), 3.67–3.56 (m, 1H), 3.55–3.45 (m, 1H), 2.92–2.79 (m, 1H), 2.79–2.65 (m, 1H), 2.38–2.15 (m, 4H), 2.15–1.85 (m, 2H), 1.85–1.72 (m, 2H), 1.34–1.22 (m, 2H), 0.95 (t, $J = 6.6$ Hz, 3H); ^{13}C NMR (75 MHz, CDCl_3) δ 207.5, 141.5, 128.5, 126.0, 75.9, 75.3, 48.4, 48.1, 45.7, 38.1, 31.7, 24.6, 23.3, 22.2; IR (thin film): 2956, 2927, 1720, 1368, 1055, 700 cm^{-1} ; HRMS (ES/MeOH): m/z calcd for $\text{C}_{17}\text{H}_{24}\text{NaO}_2$ ($\text{M} + \text{Na}$) $^+$ 283.1674, found 283.1641.



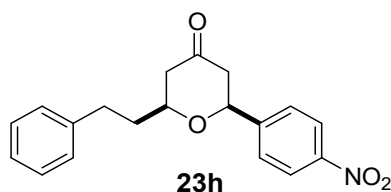
2-Isopropyl-6-phenethyltetrahydropyran-4-one (23e). Light yellow oil: $R_f = 0.46$ (20% EtOAc/hexanes); ^1H NMR (300 MHz, CDCl_3) δ 7.32–7.23 (m, 2H), 7.22–7.14 (m, 3H), 3.54–3.43 (m, 1H), 3.23 (ddd, $J = 11.4, 6.6, 2.4$ Hz, 1H), 2.91–2.79 (m, 1H), 2.79–2.66 (m, 1H), 2.43–2.32 (m, 1H), 2.31–2.16 (m, 3H), 2.05–1.90 (m, 1H), 1.88–1.71 (m, 2H), 1.05 (d, $J = 6.6$ Hz, 3H), 0.94 (d, $J = 6.9$ Hz, 3H); ^{13}C NMR (75 MHz, CDCl_3) δ 208.1, 141.6, 128.5, 126.0, 81.9, 75.6, 48.0, 45.3, 38.1, 33.6, 31.6, 18.5, 18.3; IR (thin film): 3026, 2961, 1719, 1456, 1062, 700 cm^{-1} ; HRMS (ES/MeOH): m/z calcd for $\text{C}_{16}\text{H}_{23}\text{O}_2$ ($\text{M} + \text{H}$) $^+$ 247.1698, found 247.1630.



2-Cyclohexyl-6-phenethyltetrahydropyran-4-one (23f). Light yellow oil: $R_f = 0.48$ (20% EtOAc/hexanes); $^1\text{H NMR}$ (300 MHz, CDCl_3) δ 7.34–7.24 (m, 2H), 7.24–7.15 (m, 3H), 3.53–3.41 (m, 1H), 3.26 (ddd, $J = 11.4, 6.9, 2.4$ Hz, 1H), 2.91–2.67 (m, 2H), 2.46–2.33 (m, 1H), 2.32–2.17 (m, 3H), 2.14–1.90 (m, 2H), 1.85–1.65 (m, 5H), 1.59–1.39 (m, 1H), 1.38–0.95 (m, 5H); $^{13}\text{C NMR}$ (75 MHz, CDCl_3) δ 208.2, 141.6, 128.6, 128.5, 126.0, 81.2, 75.6, 48.2, 45.3, 43.4, 38.1, 31.6, 29.0, 28.6, 26.5, 26.1, 26.0; IR (thin film): 2925, 2852, 1718, 1451, 1059, 700 cm^{-1} ; HRMS (ES/MeOH): m/z calcd for $\text{C}_{19}\text{H}_{26}\text{NaO}_2$ ($\text{M} + \text{Na}$) $^+$ 309.1830, found 309.1814.

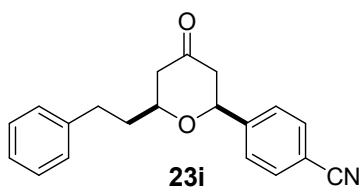


2-Phenethyl-6-phenyltetrahydropyran-4-one (23g). Light yellow oil: $R_f = 0.40$ (20% EtOAc/hexanes); $^1\text{H NMR}$ (300 MHz, CDCl_3) δ 7.38 (d, $J = 4.2$ Hz, 4H); 7.36–7.23 (m, 3H), 7.22–7.14 (m, 3H), 4.60 (dd, $J = 11.4, 3.0$ Hz, 1H), 3.78–3.67 (m, 1H), 2.92–2.70 (m, 2H), 2.68–2.51 (m, 2H), 2.50–2.35 (m, 2H), 2.16–2.01 (m, 1H), 1.96–1.81 (m, 1H); $^{13}\text{C NMR}$ (75 MHz, CDCl_3) δ 206.6, 141.4, 141.0, 128.6, 128.5, 128.0, 126.0, 125.6, 78.4, 76.2, 49.5, 47.7, 37.9, 31.5; IR (thin film): 3027, 2927, 1718, 1454, 1058, 699 cm^{-1} ; HRMS (ES/MeOH): m/z calcd for $\text{C}_{19}\text{H}_{20}\text{NaO}_2$ ($\text{M} + \text{Na}$) $^+$ 303.1361, found 303.1329.

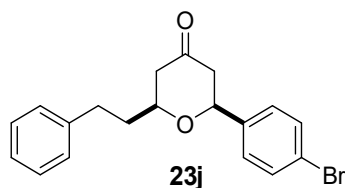


2-(4-Nitrophenyl)-6-phenethyltetrahydropyran-4-one (23h). Light yellow oil: $R_f = 0.30$ (20% EtOAc/hexanes); $^1\text{H NMR}$ (300 MHz, CDCl_3) δ 8.25 (app d, $J = 8.7$ Hz, 2H); 7.55 (d, $J = 8.7$ Hz, 2H); 7.32–7.25 (m, 2H), 7.24–7.16 (m, 3H), 4.73 (dd, $J = 11.7, 2.7$ Hz, 1H), 3.83–3.72 (m, 1H), 2.94–2.73 (m, 2H), 2.72–2.63 (m, 1H), 2.54–2.40 (m,

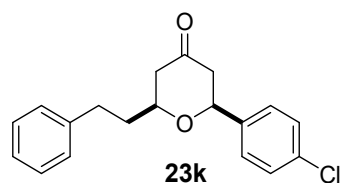
3H), 2.20–2.06 (m, 1H), 2.02–1.88 (m, 1H); ^{13}C NMR (75 MHz, CDCl_3) δ 205.2, 148.1, 147.7, 141.2, 128.6, 128.5, 126.3, 126.2, 124.0, 77.4, 76.5, 49.2, 47.6, 37.8, 31.6; IR (thin film): 2927, 1719, 1519, 1345, 749, 697 cm^{-1} ; HRMS (ES/MeOH): m/z calcd for $\text{C}_{19}\text{H}_{20}\text{NO}_4$ ($\text{M} + \text{H}$) $^+$ 326.1392, found 326.1295.



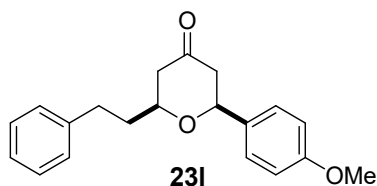
2-(4-Cyanophenyl)-6-phenethyltetrahydropyran-4-one (23i). Light yellow oil: $R_f = 0.20$ (20% EtOAc/hexanes); ^1H NMR (300 MHz, CDCl_3) δ 7.68 (d, $J = 5.1$ Hz, 2H); 7.49 (d, $J = 5.1$ Hz, 2H); 7.30–7.26 (m, 2H), 7.21–7.17 (m, 3H), 4.67 (dd, $J = 7.2, 1.5$ Hz, 1H), 3.78–3.73 (m, 1H), 2.89–2.83 (m, 1H), 2.81–2.75 (m, 1H), 2.66–2.63 (m, 1H), 2.49–2.39 (m, 3H), 2.15–2.07 (m, 1H), 1.96–1.89 (m, 1H); ^{13}C NMR (75 MHz, CDCl_3) δ 205.5, 146.1, 141.2, 132.5, 128.5, 128.4, 126.2, 126.1, 118.7, 111.8, 77.4, 76.3, 49.1, 47.5, 37.8, 31.5; IR (thin film): 3026, 2926, 2228, 1721, 1344, 701 cm^{-1} ; HRMS (ES/MeOH): m/z calcd for $\text{C}_{20}\text{H}_{19}\text{NNaO}_2$ ($\text{M} + \text{Na}$) $^+$ 328.1313, found 328.1311.



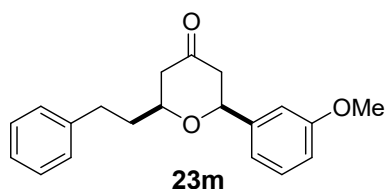
2-(4-Bromophenyl)-6-phenethyltetrahydropyran-4-one (23j). Light yellow oil: $R_f = 0.40$ (20% EtOAc/hexanes); ^1H NMR (300 MHz, CDCl_3) δ 7.52 (d, $J = 8.7$ Hz, 2H), 7.32–7.23 (m, 4H), 7.23–7.14 (m, 3H), 4.58 (dd, $J = 11.4, 2.7$ Hz, 1H), 3.78–3.67 (m, 1H), 2.92–2.71 (m, 2H), 2.66–2.57 (m, 1H), 2.54–2.37 (m, 3H), 2.16–2.02 (m, 1H), 1.97–1.83 (m, 1H); ^{13}C NMR (75 MHz, CDCl_3) δ 206.3, 141.3, 140.0, 131.8, 128.6, 128.5, 127.4, 126.1, 121.9, 77.8, 76.3, 49.4, 47.7, 37.9, 31.5; IR (thin film): 3026, 2927, 1718, 1489, 1070, 700 cm^{-1} ; HRMS (ES/MeOH): m/z calcd for $\text{C}_{19}\text{H}_{20}\text{BrO}_2$ ($\text{M} + \text{H}$) $^+$ 359.0647, found 359.0468.



2-(4-Chlorophenyl)-6-phenethyltetrahydropyran-4-one (23k). Light yellow oil: $R_f = 0.40$ (20% EtOAc/hexanes); $^1\text{H NMR}$ (300 MHz, CDCl_3) δ 7.39–7.24 (m, 6H), 7.23–7.15 (m, 3H), 4.59 (dd, $J = 11.7, 3.0$ Hz, 1H), 3.78–3.67 (m, 1H), 2.91–2.71 (m, 2H), 2.66–2.57 (m, 1H), 2.55–2.37 (m, 3H), 2.17–2.02 (m, 1H), 1.98–1.83 (m, 1H); $^{13}\text{C NMR}$ (75 MHz, CDCl_3) δ 206.4, 141.3, 139.5, 133.8, 128.8, 128.6, 128.5, 127.0, 126.1, 77.8, 76.3, 49.4, 47.7, 37.9, 31.5; IR (thin film): 3026, 2927, 1717, 1490, 1088, 700 cm^{-1} ; HRMS (ES/MeOH): m/z calcd for $\text{C}_{19}\text{H}_{19}\text{ClNaO}_2$ ($\text{M} + \text{Na}$) $^+$ 337.0971, found 337.0925.

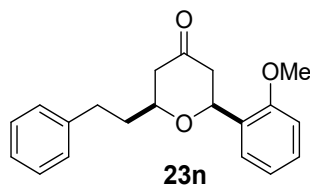


2-(4-Methoxyphenyl)-6-phenethyltetrahydropyran-4-one (23l). Light yellow oil: $R_f = 0.32$ (20% EtOAc/hexanes); $^1\text{H NMR}$ (300 MHz, CDCl_3) δ 7.36–7.24 (m, 4H), 7.23–7.15 (m, 3H), 6.93 (d, $J = 8.7$ Hz, 2H), 4.57 (dd, $J = 9.6, 4.5$ Hz, 1H), 3.82 (s, 3H), 3.77–3.67 (m, 1H), 2.91–2.70 (m, 2H), 2.67–2.53 (m, 2H), 2.50–2.33 (m, 2H), 2.16–2.00 (m, 1H), 1.96–1.82 (m, 1H); $^{13}\text{C NMR}$ (75 MHz, CDCl_3) δ 206.9, 159.4, 141.5, 133.2, 128.5, 127.1, 126.1, 114.1, 78.3, 76.2, 55.4, 49.5, 47.8, 38.0, 31.6; IR (thin film): 3026, 2931, 1717, 1493, 1052, 697 cm^{-1} ; HRMS (ES/MeOH): m/z calcd for $\text{C}_{20}\text{H}_{22}\text{NaO}_3$ ($\text{M} + \text{Na}$) $^+$ 333.1467, found 333.1442.

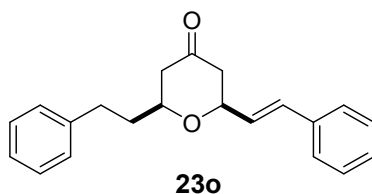


2-(3-Methoxyphenyl)-6-phenethyltetrahydropyran-4-one (23m). Light yellow oil: $R_f = 0.34$ (20% EtOAc/hexanes); $^1\text{H NMR}$ (300 MHz, CDCl_3) δ 7.35–7.23 (m, 3H), 7.23–7.14 (m, 3H), 7.00–6.92 (m, 2H), 6.90–6.82 (m, 1H), 4.60 (dd, $J = 11.1, 3.0$ Hz, 1H), 3.83 (s, 3H), 3.78–3.67 (m, 1H), 2.93–2.71 (m, 2H), 2.69–2.37 (m, 4H), 2.18–2.01

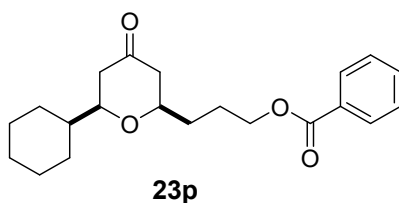
(m, 1H), 1.97–1.81 (m, 1H); ^{13}C NMR (75 MHz, CDCl_3) δ 206.7, 159.9, 142.6, 141.4, 129.7, 128.5, 126.0, 117.9, 113.2, 111.5, 78.3, 76.2, 55.3, 49.5, 47.7, 37.9, 31.5; IR (thin film): 2933, 2859, 1718, 1490, 1046, 698 cm^{-1} ; HRMS (ES/MeOH): m/z calcd for $\text{C}_{20}\text{H}_{22}\text{NaO}_3$ ($\text{M} + \text{Na}$) $^+$ 333.1467, found 333.1430.



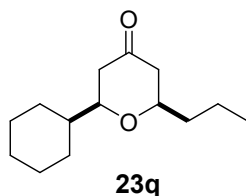
2-(2-Methoxyphenyl)-6-phenethyltetrahydropyran-4-one (23n). Light yellow oil: $R_f = 0.37$ (20% EtOAc/hexanes); ^1H NMR (300 MHz, CDCl_3) δ 7.58 (d, $J = 7.5$ Hz, 1H), 7.32–7.22 (m, 3H), 7.22–7.13 (m, 3H), 7.03 (t, $J = 7.5$ Hz, 1H), 6.87 (d, $J = 8.1$ Hz, 1H), 4.96 (dd, $J = 11.4, 2.1$ Hz, 1H), 3.81 (s, 3H), 3.80–3.70 (m, 1H), 2.91–2.69 (m, 3H), 2.48–2.28 (m, 3H), 2.16–2.01 (m, 1H), 1.96–1.82 (m, 1H); ^{13}C NMR (75 MHz, CDCl_3) δ 207.5, 155.5, 141.6, 129.7, 128.7, 128.6, 128.5, 126.0, 120.9, 110.3, 76.1, 73.3, 55.3, 48.3, 47.8, 38.1, 31.5; IR (thin film): 3026, 2928, 1717, 1494, 1246, 754 cm^{-1} ; HRMS (ES/MeOH): m/z calcd for $\text{C}_{20}\text{H}_{22}\text{NaO}_3$ ($\text{M} + \text{Na}$) $^+$ 333.1467, found 333.1435.



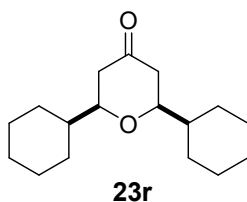
2-Phenethyl-6-styryltetrahydropyran-4-one (23o). Light yellow oil: $R_f = 0.38$ (20% EtOAc/hexanes); ^1H NMR (300 MHz, CDCl_3) δ 7.43–7.33 (m, 4H), 7.31–7.19 (m, 6H), 6.65 (d, $J = 15.9$ Hz, 1H), 6.26 (d, $J = 15.9, 5.7$ Hz, 1H), 4.26–4.20 (m, 1H), 3.69–3.61 (m, 1H), 2.90–2.71 (m, 2H), 2.54–2.28 (m, 4H), 2.12–2.00 (m, 1H), 1.91–1.79 (m, 1H); ^{13}C NMR (75 MHz, CDCl_3) δ 206.5, 141.1, 136.4, 131.3, 128.7, 128.5, 128.4, 128.1, 126.7, 126.1, 77.3, 76.0, 47.9, 47.8, 37.9, 31.6; IR (thin film): 3026, 2927, 1718, 1495, 1054, 747 cm^{-1} ; HRMS (ES/MeOH): m/z calcd for $\text{C}_{21}\text{H}_{22}\text{NaO}_2$ ($\text{M} + \text{Na}$) $^+$ 329.1517, found 329.1511.



2-(3-Benzoyloxy)propyl-6-cyclohexyltetrahydropyran-4-one (23p). Colorless oil: $R_f = 0.44$ (20% EtOAc/hexanes); $^1\text{H NMR}$ (300 MHz, CDCl_3) δ 8.08–8.01 (m, 2H), 7.57 (app t, $J = 7.2$ Hz, 1H), 7.45 (app t, $J = 7.2$ Hz, 2H), 4.44–4.39 (m, 2H), 3.63–3.52 (m, 1H), 3.29 (ddd, $J = 11.4, 6.6, 2.4$ Hz, 1H), 2.45–2.33 (m, 2H), 2.31–2.18 (m, 2H), 2.10–1.59 (m, 10H), 1.56–1.41 (m, 1H), 1.33–1.10 (m, 2H), 1.10–0.93 (m, 2H); $^{13}\text{C NMR}$ (75 MHz, CDCl_3) 208.1, 166.7, 133.0, 130.5, 129.7, 128.5, 81.4, 76.6, 64.8, 48.3, 45.4, 43.3, 33.1, 28.9, 28.6, 26.5, 26.1, 26.0, 25.1; IR (thin film): 2926, 2852, 1718, 1451, 1274, 1113 cm^{-1} ; HRMS (ES/MeOH): m/z calcd for $\text{C}_{21}\text{H}_{28}\text{NaO}_4$ ($\text{M} + \text{Na}$) $^+$ 367.1885, found 367.1877.

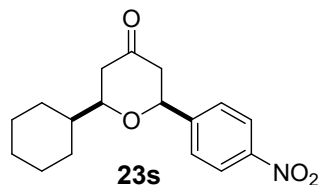


2-Cyclohexyl-6-propyltetrahydropyran-4-one (23q). Colorless oil: $R_f = 0.53$ (20% EtOAc/hexanes); $^1\text{H NMR}$ (300 MHz, CDCl_3) δ 3.52–3.48 (m, 1H), 3.26 (ddd, $J = 6.9, 4.5, 1.5$ Hz, 1H), 2.36 (dd, $J = 15.3, 8.4$ Hz, 2H), 2.21 (dd, $J = 15.3, 2.7$ Hz, 2H), 2.01–1.98 (m, 1H), 1.76–1.63 (m, 5H), 1.53–1.38 (m, 5H), 1.25–1.14 (m, 2H), 1.04–0.98 (m, 2H), 0.93 (t, $J = 4.5$ Hz, 3H); $^{13}\text{C NMR}$ (75 MHz, CDCl_3) δ 208.7, 81.3, 76.8, 48.4, 45.6, 43.3, 38.6, 29.0, 28.5, 26.5, 26.1, 26.0, 18.8, 14.0; IR (thin film): 2927, 2853, 1720, 1450, 1158, 846 cm^{-1} ; HRMS (ES/MeOH): m/z calcd for $\text{C}_{14}\text{H}_{24}\text{NaO}_2$ ($\text{M} + \text{Na}$) $^+$ 247.1674, found 247.1677.

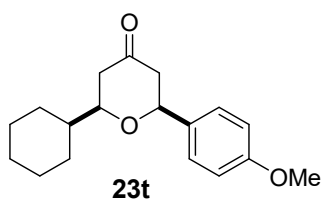


2,6-Dicyclohexyltetrahydropyran-4-one (23r). Light yellow solid: $R_f = 0.55$ (20% EtOAc/hexanes); mp 69.5–70.6 $^\circ\text{C}$; $^1\text{H NMR}$ (300 MHz, CDCl_3) δ 3.22 (ddd, $J = 11.4,$

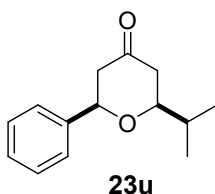
6.9, 2.1 Hz, 2H), 2.39 (dd, $J = 14.4, 1.5$ Hz, 2H), 2.28–2.15 (m, 2H), 2.06–1.93 (m, 2H), 1.84–1.58 (m, 8H), 1.54–1.39 (m, 2H), 1.34–1.10 (m, 6H), 1.10–0.92 (m, 4H); ^{13}C NMR (75 MHz, CDCl_3) δ 209.3, 81.3, 45.7, 43.4, 28.9, 28.6, 26.5, 26.1, 26.0; IR (thin film): 2925, 2852, 1719, 1450, 1271, 1066 cm^{-1} ; HRMS (ES/MeOH): m/z calcd for $\text{C}_{17}\text{H}_{28}\text{NaO}_2$ ($\text{M} + \text{Na}$) $^+$ 287.1987, found 287.1947.



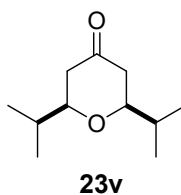
2-Cyclohexyl-6-(4-nitrophenyl)tetrahydropyran-4-one (23s). Light yellow solid: $R_f = 0.36$ (20% EtOAc/hexanes); mp 118.2–119.8 $^{\circ}\text{C}$; ^1H NMR (500 MHz, CDCl_3) δ 8.24 (d, $J = 5.1$ Hz, 2H), 7.55 (d, $J = 5.1$ Hz, 2H), 4.72, $J = 6.9, 1.2$ Hz, 1H), 3.58–3.54 (m, 1H), 2.69–2.67 (m, 1H), 2.52–2.40 (m, 3H), 1.99–1.97 (m, 1H), 1.80–1.62 (m, 5H), 1.33–1.06 (m, 5H); ^{13}C NMR (125 MHz, CDCl_3) δ 206.3, 148.4, 147.5, 126.3, 123.9, 81.6, 77.4, 49.5, 45.0, 43.2, 28.7, 28.5, 26.5, 26.1, 26.0; IR (thin film): 2927, 2853, 1721, 1520, 1346, 853 cm^{-1} ; HRMS (ES/MeOH): m/z calcd for $\text{C}_{17}\text{H}_{22}\text{NO}_4$ ($\text{M} + \text{H}$) $^+$ 304.1549, found 304.1450.



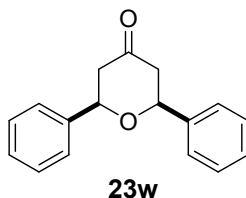
2-Cyclohexyl-6-(4-methoxyphenyl)tetrahydropyran-4-one (23t). Colorless oil: $R_f = 0.47$ (20% EtOAc/hexanes); ^1H NMR (300 MHz, CDCl_3) δ 7.29 (d, $J = 5.1$ Hz, 2H), 6.90 (d, $J = 5.1$ Hz, 2H), 4.55, $J = 6.9, 1.5$ Hz, 1H), 3.80 (s, 3H), 3.52–3.48 (m, 1H), 2.62–2.50 (m, 2H), 2.46–2.35 (m, 2H), 1.97–1.95 (m, 1H), 1.77–1.56 (m, 5H), 1.27–1.04 (m, 5H); ^{13}C NMR (75 MHz, CDCl_3) δ 208.0, 159.3, 133.4, 126.9, 114.0, 81.5, 78.2, 55.4, 49.8, 45.1, 43.2, 28.8, 28.4, 26.5, 26.1; IR (thin film): 2926, 2852, 1719, 1514, 1249, 827 cm^{-1} ; HRMS (ES/MeOH): m/z calcd for $\text{C}_{18}\text{H}_{25}\text{O}_3$ ($\text{M} + \text{H}$) $^+$ 289.1804, found 289.1800.



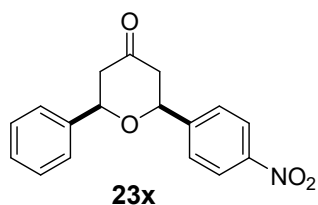
2-Isopropyl-6-phenyltetrahydropyran-4-one (23u). Light yellow oil: $R_f = 0.53$ (20% EtOAc/hexanes); $^1\text{H NMR}$ (300 MHz, CDCl_3) δ 7.40–7.27 (m, 5H), 4.63 (dd, $J = 11.4, 2.7$ Hz, 1H), 3.52 (ddd, $J = 10.8, 5.7, 3.0$ Hz, 1H), 2.69–2.60 (m, 1H), 2.58–2.33 (m, 3H), 2.00–1.87 (m, 1H), 1.04 (d, $J = 6.9$ Hz, 3H), 1.00 (d, $J = 6.9$ Hz, 3H); $^{13}\text{C NMR}$ (75 MHz, CDCl_3) δ 207.6, 141.4, 128.6, 127.9, 125.6, 82.1, 78.5, 49.9, 44.8, 33.4, 18.3, 18.1; IR (thin film): 3032, 2963, 1717, 1252, 1066, 753 cm^{-1} ; HRMS (ES/MeOH): m/z calcd for $\text{C}_{14}\text{H}_{18}\text{NaO}_2$ ($\text{M} + \text{Na}$) $^+$ 241.1204, found 241.1202.



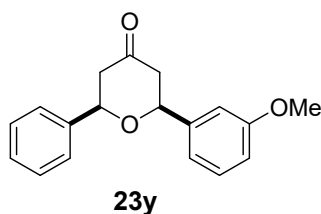
2,6-Diisopropyltetrahydropyran-4-one (23v). Light yellow oil: $R_f = 0.62$ (20% EtOAc/hexanes); $^1\text{H NMR}$ (300 MHz, CDCl_3) δ 3.22 (ddd, $J = 11.4, 6.6, 2.1$ Hz, 2H), 2.38 (dd, $J = 14.1, 1.5$ Hz, 2H), 2.25–2.16 (m, 2H), 1.83–1.72 (m, 2H), 0.99 (d, $J = 6.6$ Hz, 6H), 0.91 (d, $J = 6.6$ Hz, 6H); $^{13}\text{C NMR}$ (75 MHz, CDCl_3) δ 209.1, 81.9, 45.4, 33.6, 18.4, 18.3; IR (thin film): 2963, 1723, 1470, 1256, 1068, 887 cm^{-1} ; HRMS (ES/MeOH): m/z calcd for $\text{C}_{11}\text{H}_{20}\text{NaO}_2$ ($\text{M} + \text{Na}$) $^+$ 207.1361, found 207.1359.



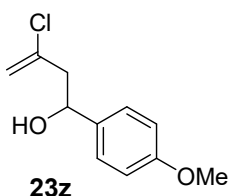
2,6-Diphenyltetrahydropyran-4-one (23w). Light yellow oil: $R_f = 0.45$ (20% EtOAc/hexanes); $^1\text{H NMR}$ (300 MHz, CDCl_3) δ 7.50–7.30 (m, 10H), 4.84 (dd, $J = 10.2, 4.2$ Hz, 2H), 2.80–2.68 (m, 4H); $^{13}\text{C NMR}$ (75 MHz, CDCl_3) δ 206.2, 140.8, 128.7, 128.2, 125.7, 79.0, 49.8; IR (thin film): 3032, 1720, 1452, 1245, 1054, 698 cm^{-1} ; HRMS (ES/MeOH): m/z calcd for $\text{C}_{17}\text{H}_{16}\text{NaO}_2$ ($\text{M} + \text{Na}$) $^+$ 275.1048, found 275.1007.



2-(4-Nitrophenyl)-6-phenyltetrahydropyran-4-one (23x). Light yellow oil: $R_f = 0.30$ (20% EtOAc/hexanes); $^1\text{H NMR}$ (300 MHz, CDCl_3) δ 8.26 (d, $J = 8.7$ Hz, 2H), 7.64 (d, $J = 8.7$ Hz, 2H), 7.50–7.30 (m, 5H), 4.97 (dd, $J = 11.7, 2.7$ Hz, 1H), 4.88 (dd, $J = 10.2, 4.2$ Hz, 1H), 2.85–2.60 (m, 4H); $^{13}\text{C NMR}$ (75 MHz, CDCl_3) δ 204.6, 147.8, 140.3, 128.9, 128.5, 126.5, 125.7, 124.0, 79.3, 77.9, 49.5, 49.3; IR (thin film): 3026, 2968, 1720, 1519, 1347, 697 cm^{-1} ; HRMS (ES/MeOH): m/z calcd for $\text{C}_{17}\text{H}_{15}\text{NNaO}_4$ ($\text{M} + \text{Na}$) $^+$ 320.0899, found 320.0895.



2-(3-Methoxyphenyl)-6-phenyltetrahydropyran-4-one (23y). Light yellow oil: $R_f = 0.38$ (20% EtOAc/hexanes); $^1\text{H NMR}$ (300 MHz, CDCl_3) δ 7.47–7.36 (m, 4H), 7.34–7.25 (m, 2H), 7.03–7.01 (m, 2H), 6.86 (dd, $J = 8.1, 2.4$ Hz, 1H), 4.86–4.79 (m, 2H), 3.82 (s, 3H), 2.76–2.64 (m, 4H); $^{13}\text{C NMR}$ (75 MHz, CDCl_3) δ 206.1, 160.0, 142.5, 140.8, 129.8, 128.7, 128.2, 125.8, 118.0, 113.3, 111.7, 79.1, 79.0, 55.4, 49.8; IR (thin film): 2962, 2836, 1717, 1267, 1041, 697 cm^{-1} ; HRMS (ES/MeOH): m/z calcd for $\text{C}_{18}\text{H}_{18}\text{NaO}_3$ ($\text{M} + \text{Na}$) $^+$ 305.1154, found 305.1150.



3-Chloro-1-(3-methoxyphenyl)but-3-en-1-ol (23z). Colorless oil: $R_f = 0.35$ (20% EtOAc/hexanes); $^1\text{H NMR}$ (300 MHz, CDCl_3) 7.28–7.24 (m, 1H), 6.96–6.94 (m, 2H), 6.85–6.81 (m, 1H), 5.29 (d, $J = 1.2$ Hz, 1H), 5.24 (d, $J = 1.2$ Hz, 1H), 5.01 (dd, $J = 8.7, 4.5$ Hz, 1H), 3.81 (s, 3H), 2.76 (dd, $J = 14.4, 8.7$ Hz, 1H), 2.65 (dd, $J = 14.4, 4.5$ Hz,

1H); ^{13}C NMR (75 MHz, CDCl_3) δ 159.8, 144.8, 138.9, 129.7, 118.1, 115.5, 113.4, 111.3, 71.1, 55.3, 49.2; IR (thin film): 3411, 2942, 1600, 1263, 1042, 882 cm^{-1} ; HRMS (ES/MeOH): m/z calcd for $\text{C}_{11}\text{H}_{13}\text{ClNaO}_2$ ($\text{M} + \text{Na}$) $^+$ 235.0502, found 235.0500.

3. ^1H and ^{13}C NMR Spectra

Figure SI1. ^1H NMR (300 MHz, CDCl_3) spectrum of 5-chloro-1-phenylhex-5-en-3-ol (**21a**).

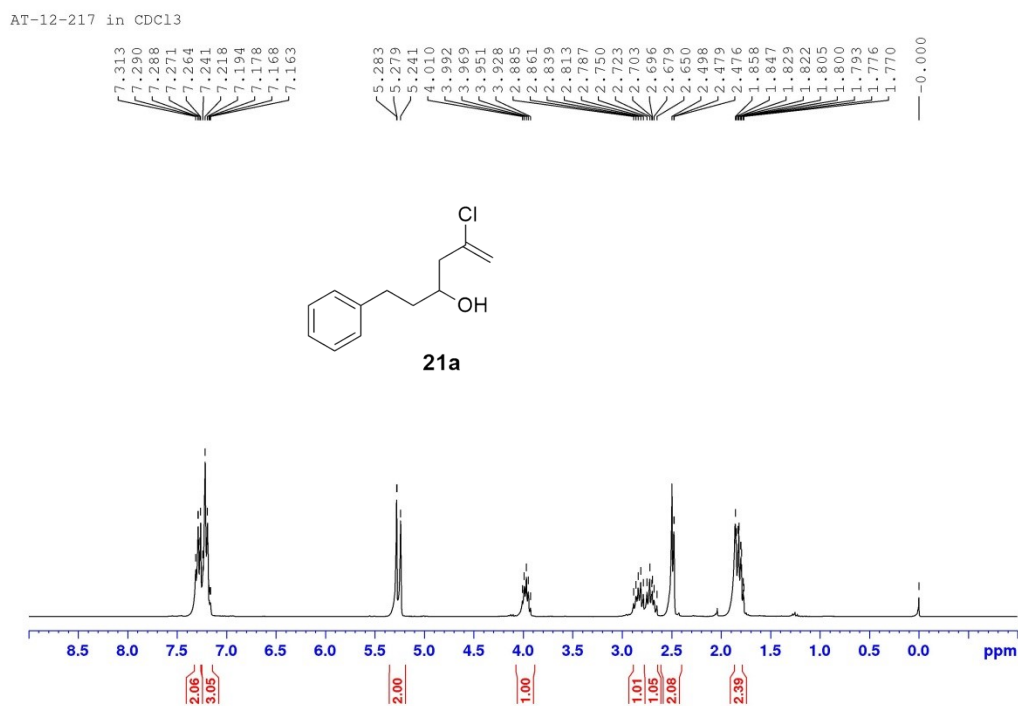


Figure SI2. ^{13}C NMR (75 MHz, CDCl_3) spectrum of 5-chloro-1-phenylhex-5-en-3-ol (**21a**).

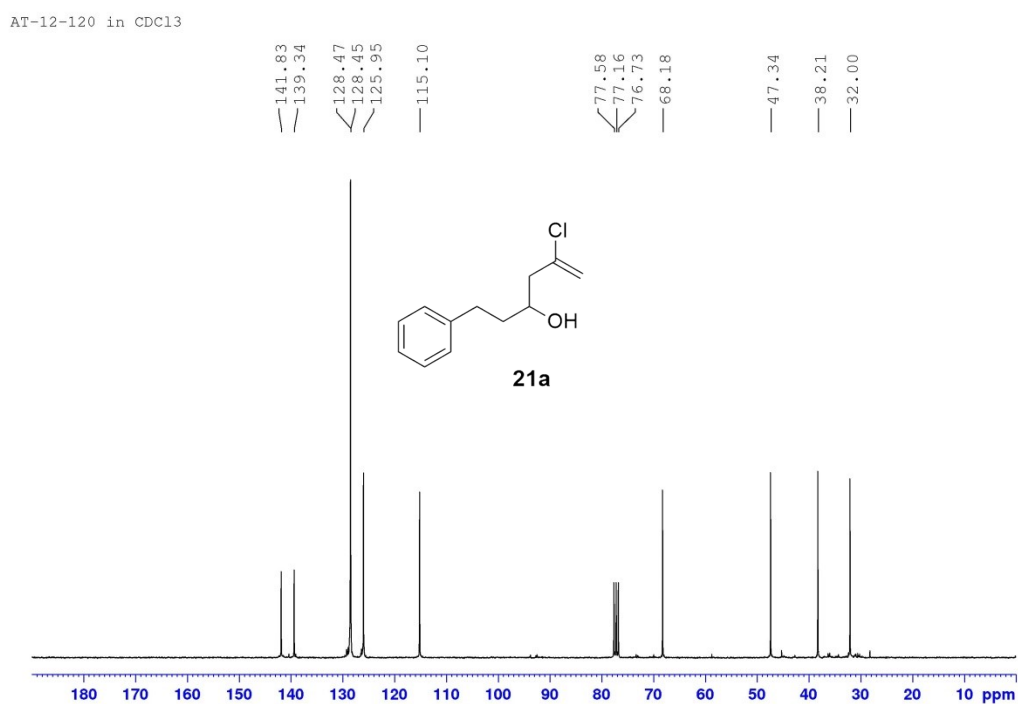


Figure SI3. ^1H NMR (300 MHz, CDCl_3) spectrum of 3-chloro-1-cyclohexyl-3-en-1-ol (**21b**).

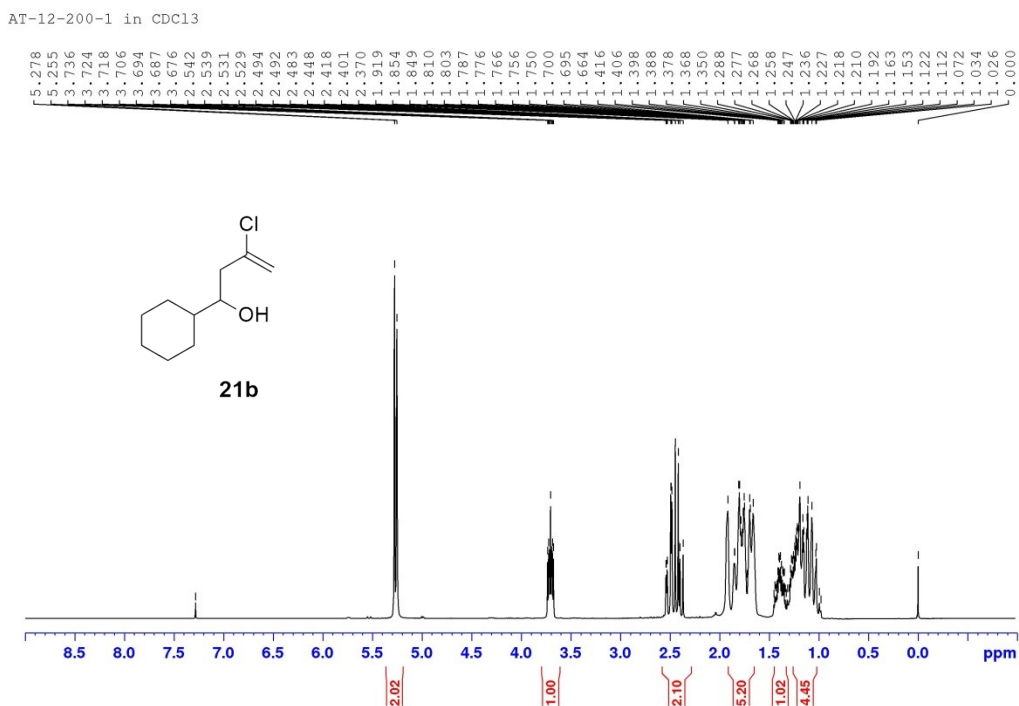


Figure SI4. ^{13}C NMR (75 MHz, CDCl_3) spectrum of 3-chloro-1-cyclohexyl-3-en-1-ol (**21b**).

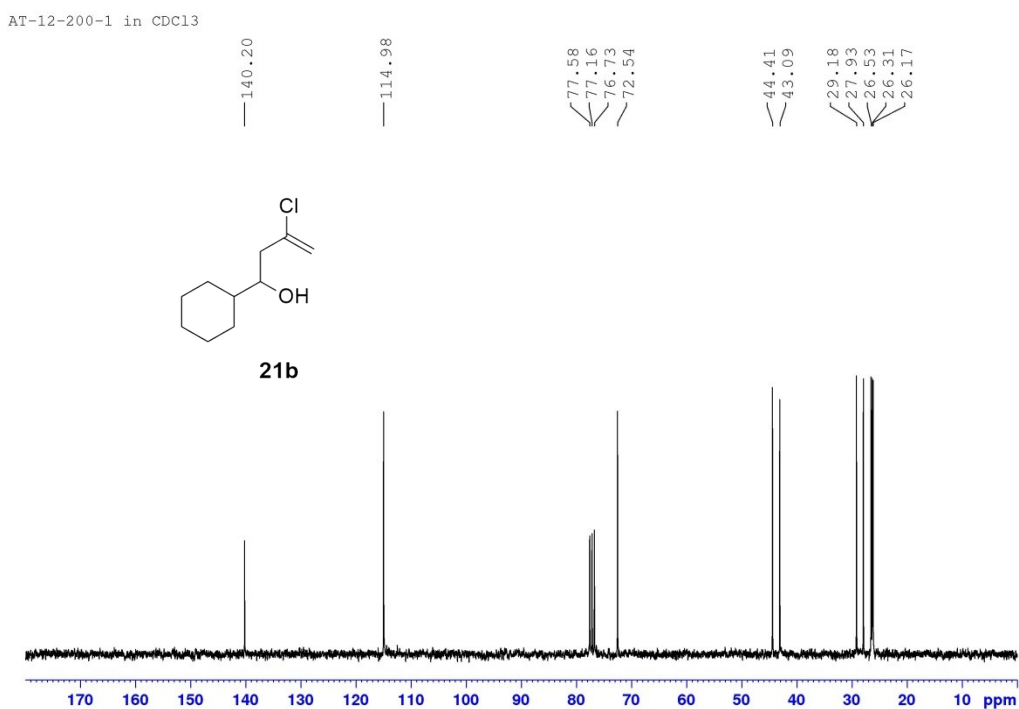


Figure SI5. ^1H NMR (300 MHz, CDCl_3) spectrum of 3-chloro-1-phenyl-3-en-1-ol (**21c**).

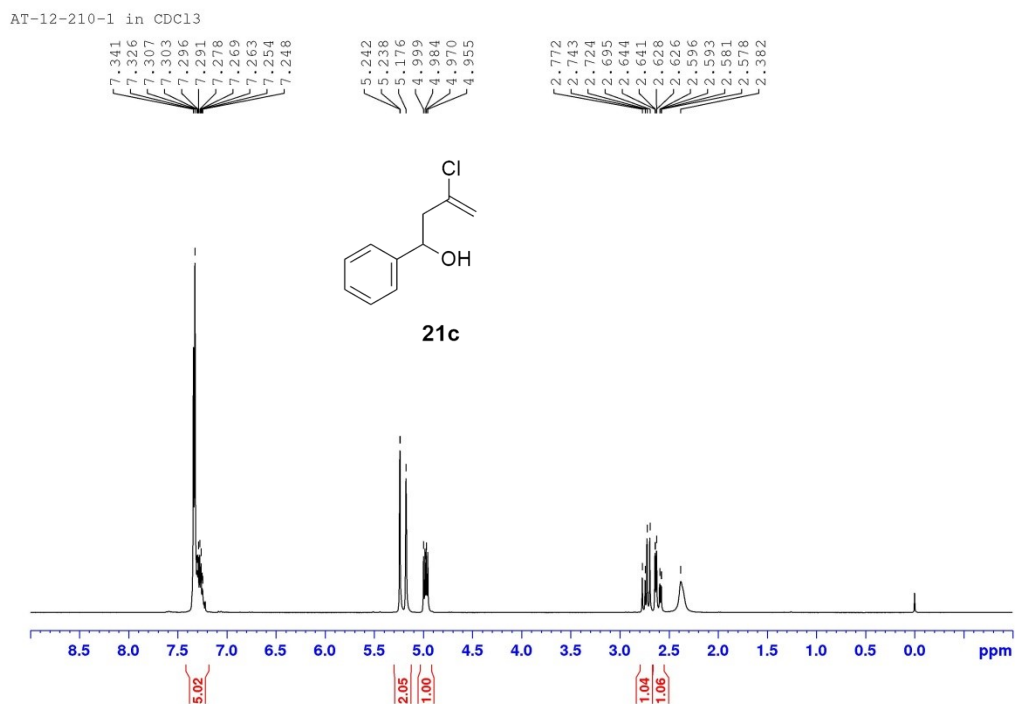


Figure SI6. ¹H NMR (75 MHz, CDCl₃) spectrum of 3-chloro-1-phenyl-3-en-1-ol (**21c**).

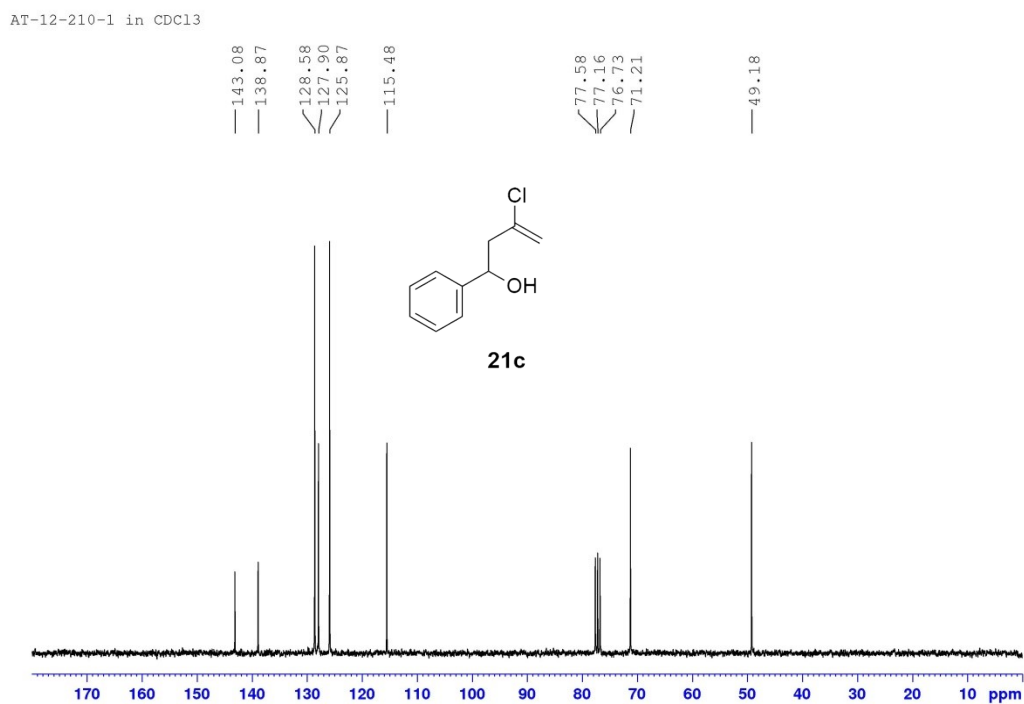


Figure SI7. ¹³C NMR (300 MHz, CDCl₃) spectrum of 2,6-diphenethyltetrahydropyran-4-one (**23a**)

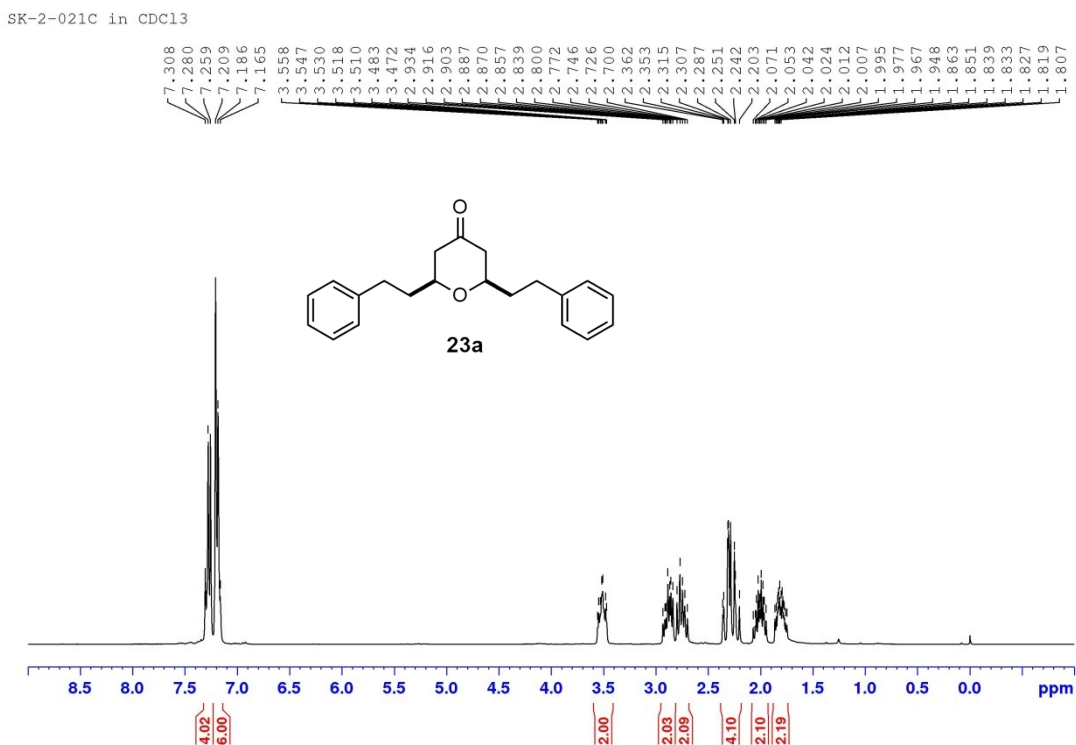


Figure SI8. ¹³C NMR (75 MHz, CDCl₃) spectrum of 2,6-diphenethyltetrahydropyran-4-one (**23a**)

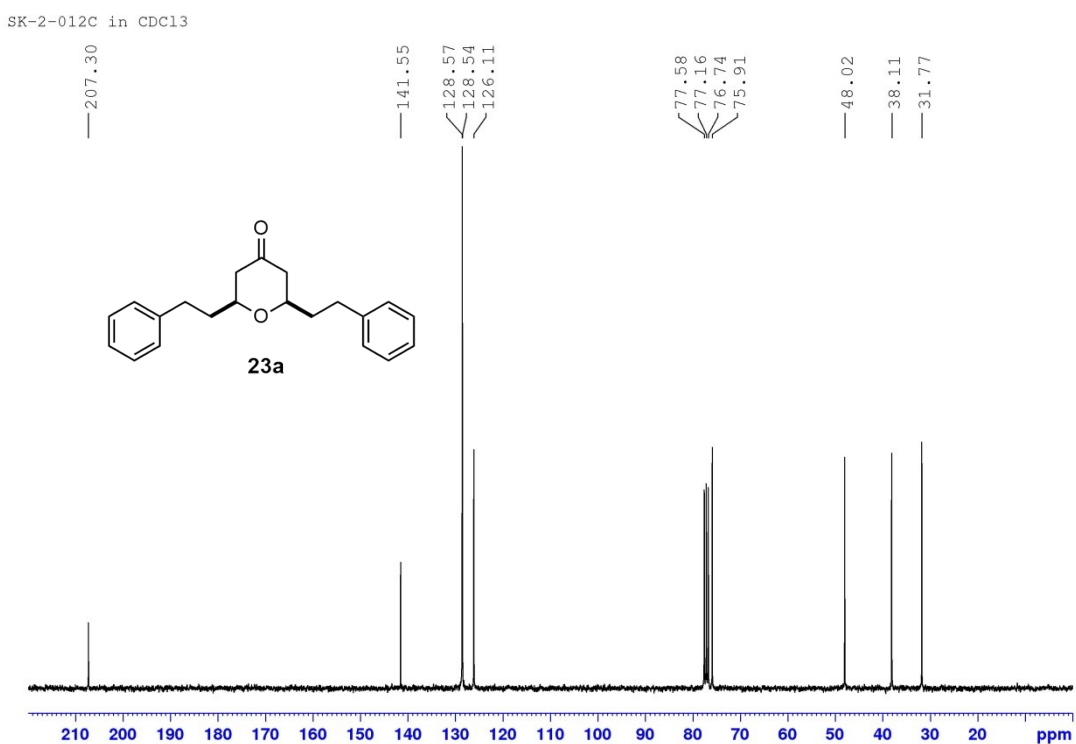


Figure SI9. ¹³C NMR (300 MHz, CDCl₃) spectrum of 4,4-dichloro-2,6-diphenethyltetrahydropyran (**24**)

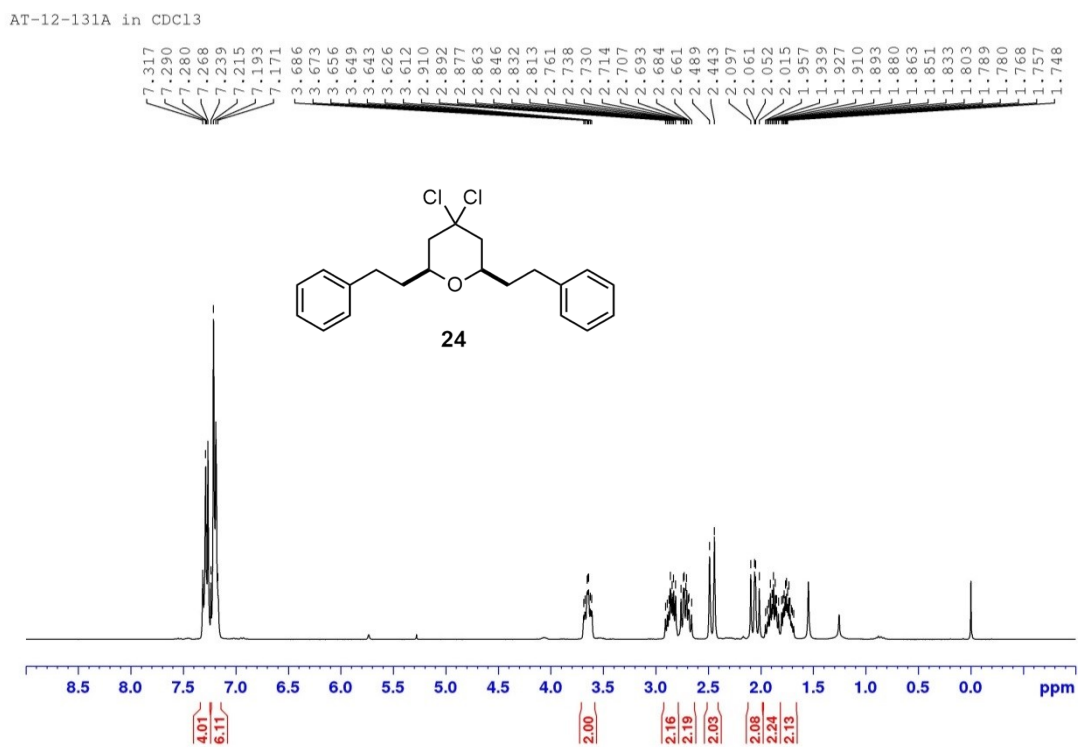


Figure SI10. ¹³C NMR (75 MHz, CDCl₃) spectrum of 4,4-dichloro-2,6-diphenethyltetrahydropyran (**24**)

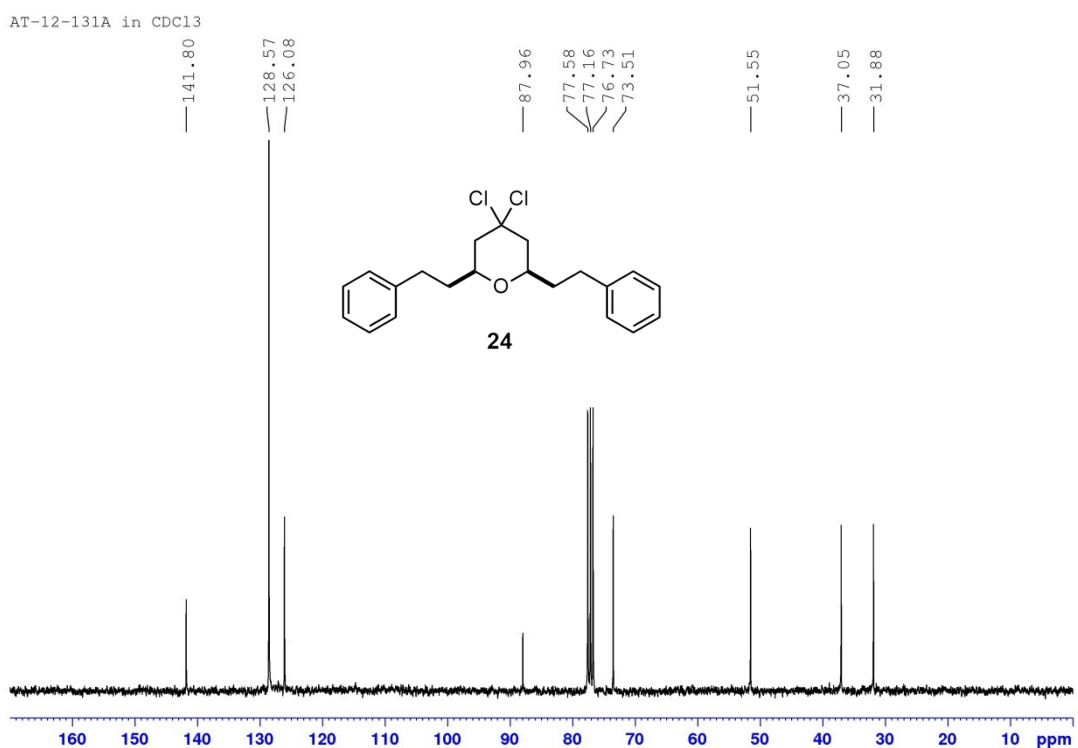


Figure SI11. ¹³C NMR (300 MHz, CDCl₃) spectrum of 2-phenethyl-6-propyltetrahydropyran-4-one (**23b**)

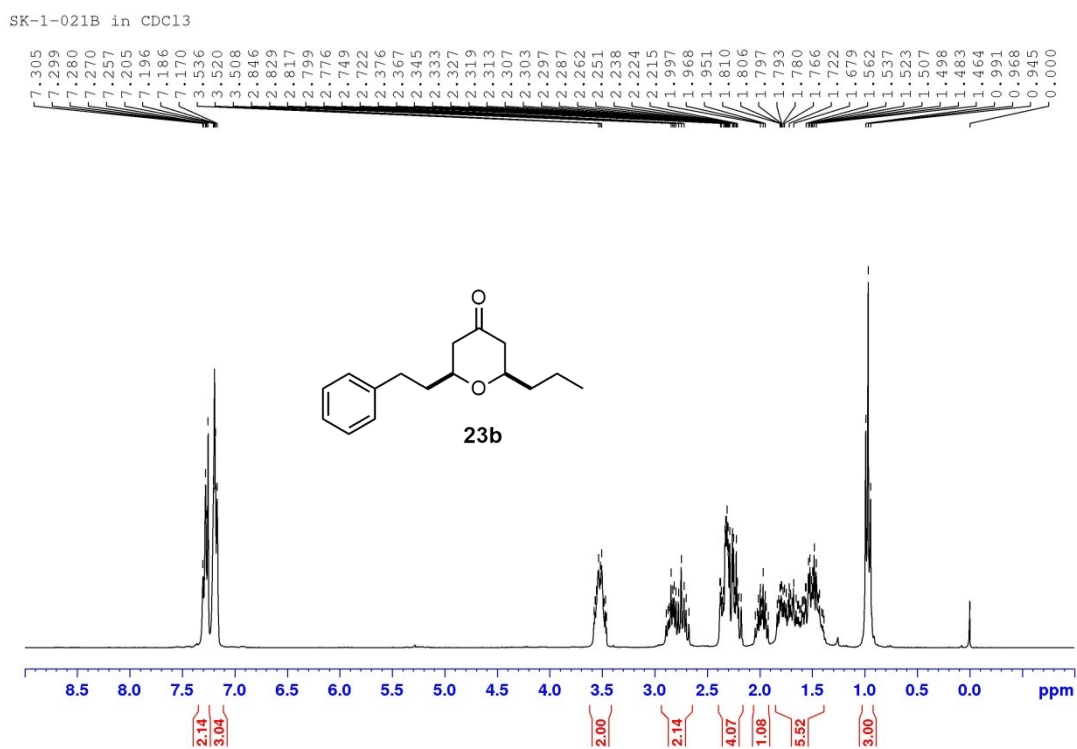


Figure SI12. ¹³C NMR (75 MHz, CDCl₃) spectrum of 2-phenethyl-6-propyltetrahydropyran-4-one (**23b**)

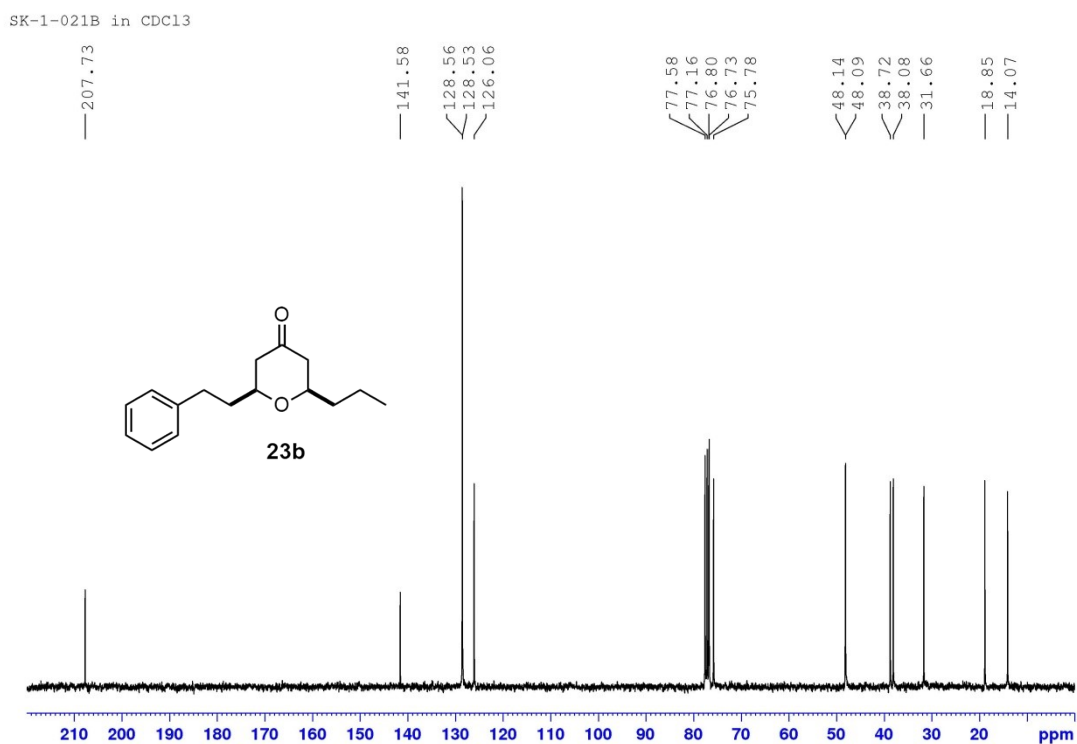


Figure SI13. ¹³C NMR (300 MHz, CDCl₃) spectrum of 2-pentyl-6-phenethyltetrahydropyran-4-one (**23c**)

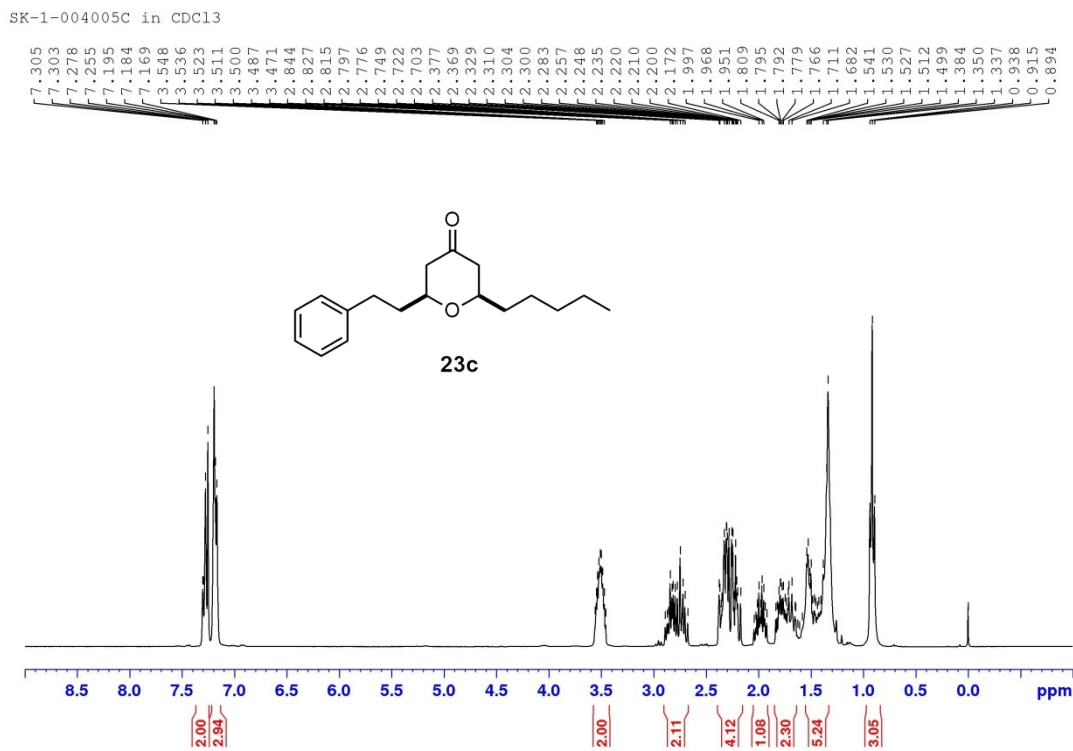


Figure SI14. ¹³C NMR (75 MHz, CDCl₃) spectrum of 2-pentyl-6-phenyltetrahydropyran-4-one (**23c**)

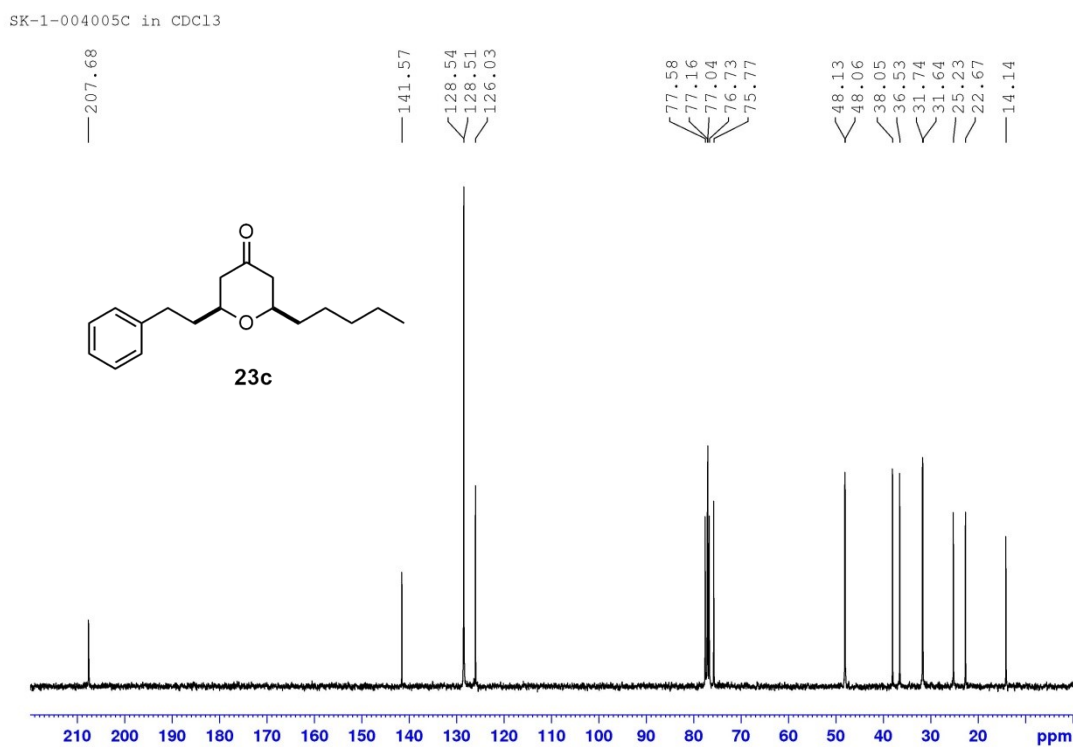


Figure SI15. ¹H NMR (300 MHz, CDCl₃) spectrum of 2-isobutyl-6-phenyltetrahydropyran-4-one (**23d**)

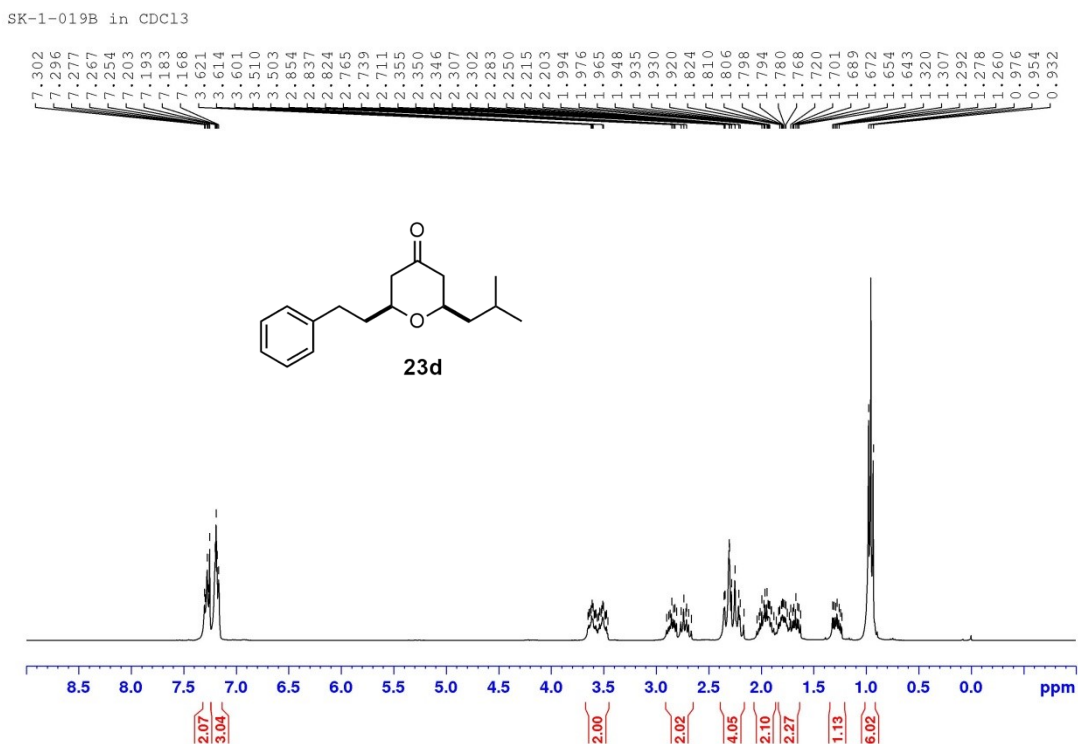


Figure SI16. ¹³C NMR (75 MHz, CDCl₃) spectrum of 2-isobutyl-6-phenethyltetrahydropyran-4-one (**23d**)

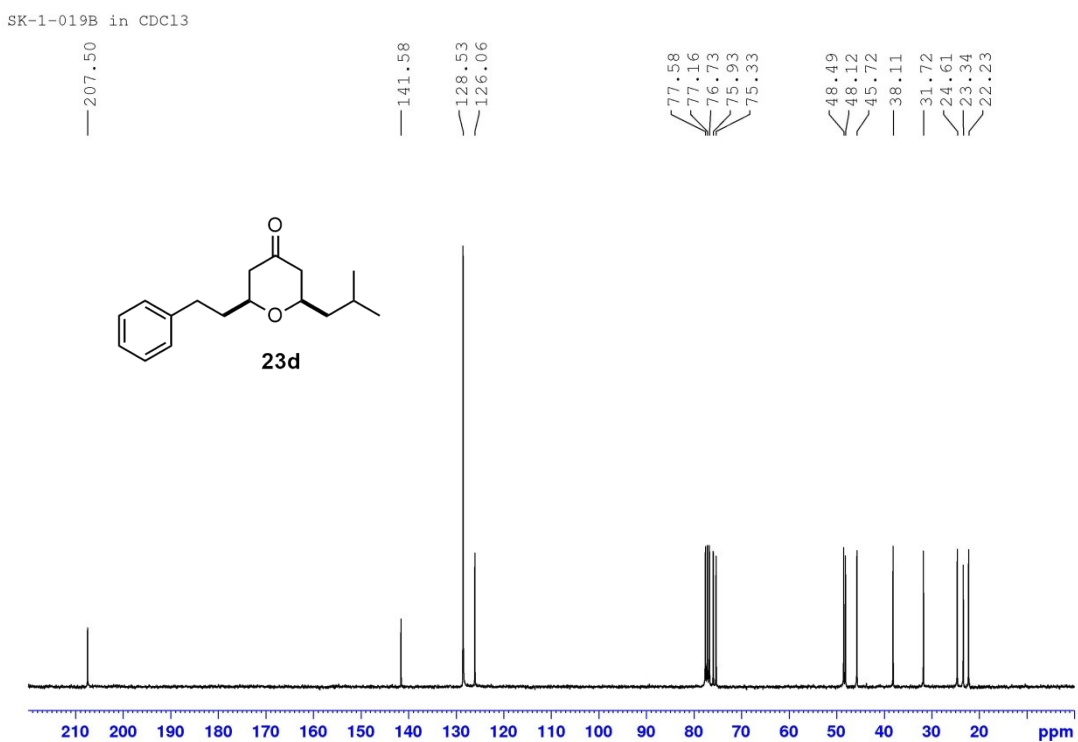


Figure SI17. ¹H NMR (300 MHz, CDCl₃) spectrum of 2-isopropyl-6-phenethyltetrahydropyran-4-one (**23e**)

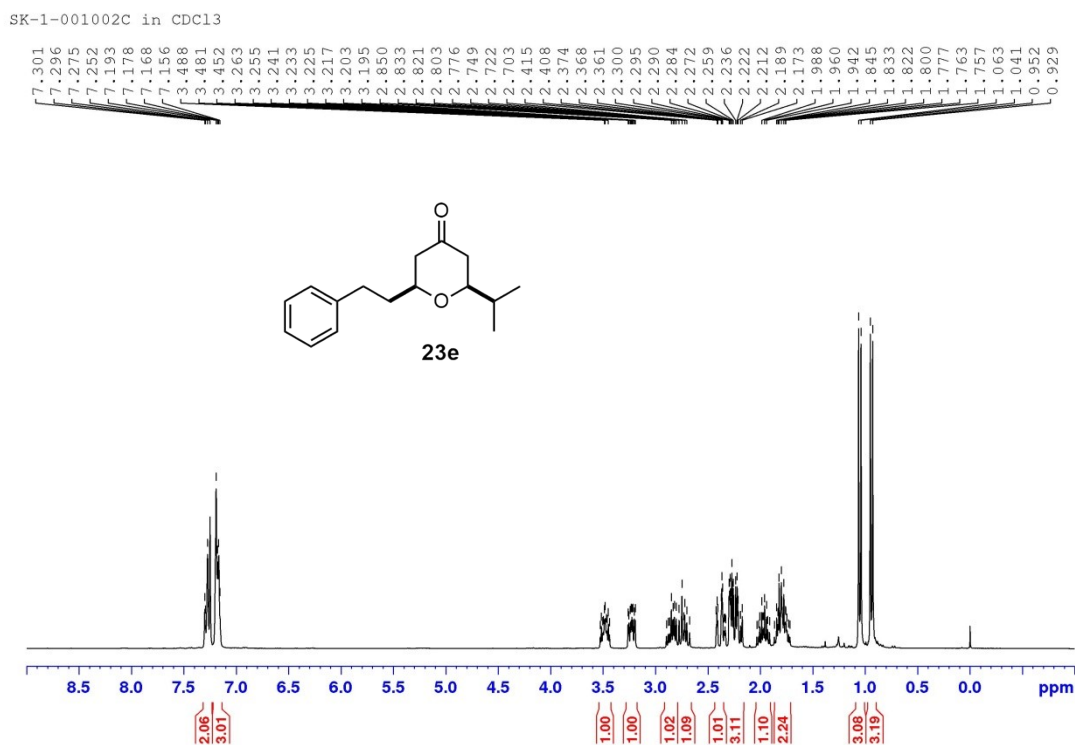


Figure SI18. ¹³C NMR (75 MHz, CDCl₃) spectrum of 2-isopropyl-6-phenethyltetrahydropyran-4-one (**23e**)

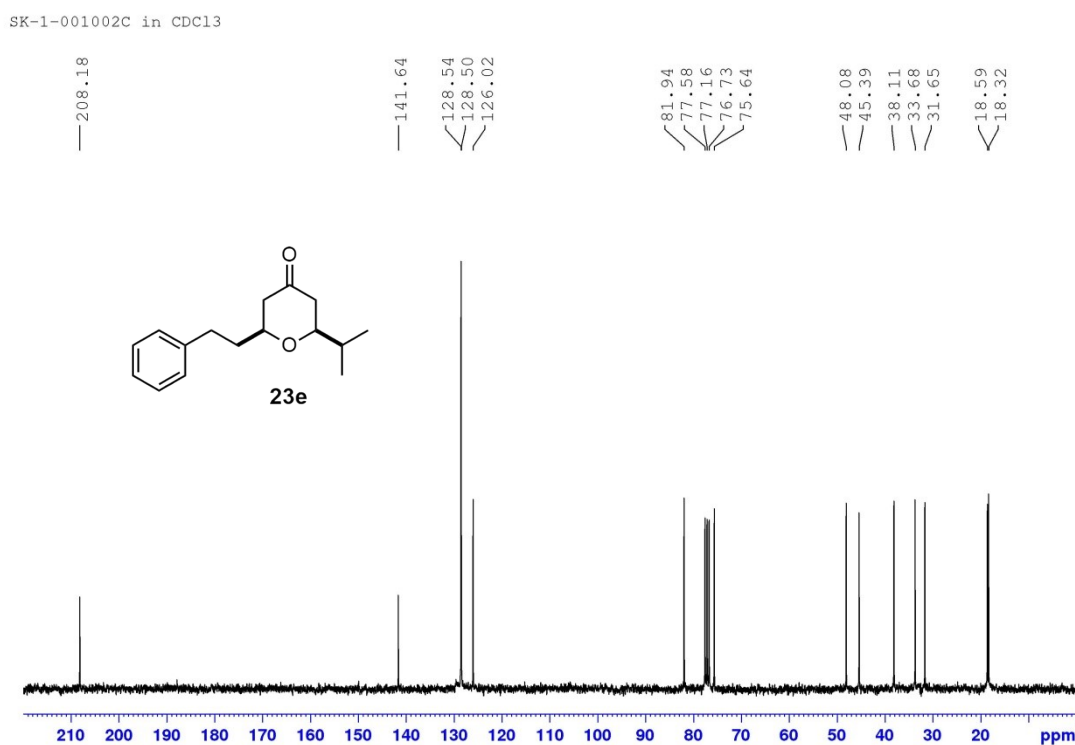


Figure SI19. ¹H NMR (300 MHz, CDCl₃) spectrum of 2-cyclohexyl-6-phenethyltetrahydropyran-4-one (**23f**)

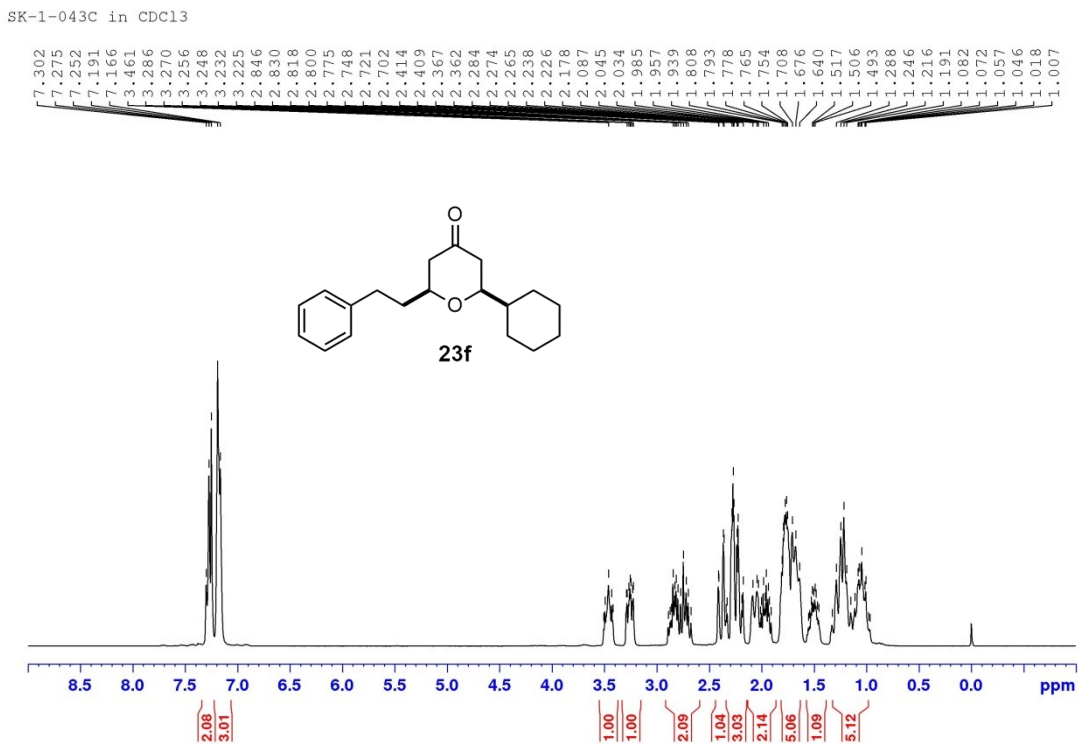


Figure SI20. ¹³C NMR (75 MHz, CDCl₃) spectrum of 2-cyclohexyl-6-phenethyltetrahydropyran-4-one (**23f**)

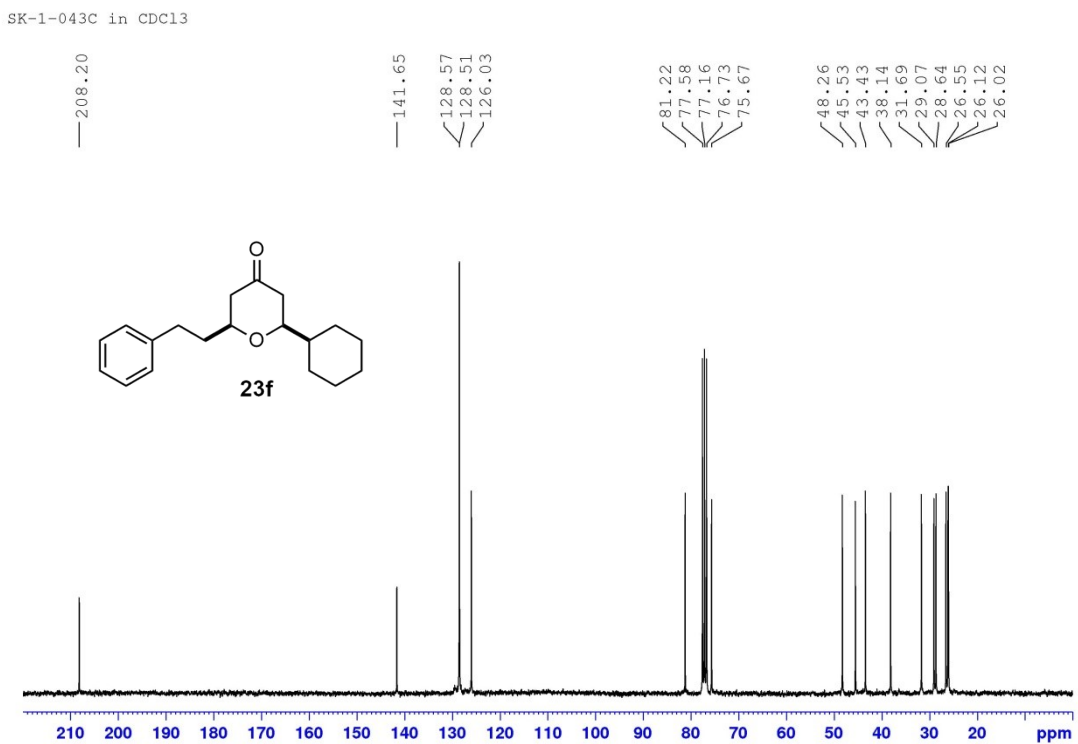


Figure SI21. ¹H NMR (300 MHz, CDCl₃) spectrum of 2-phenethyl-6-phenyltetrahydropyran-4-one (**23g**)

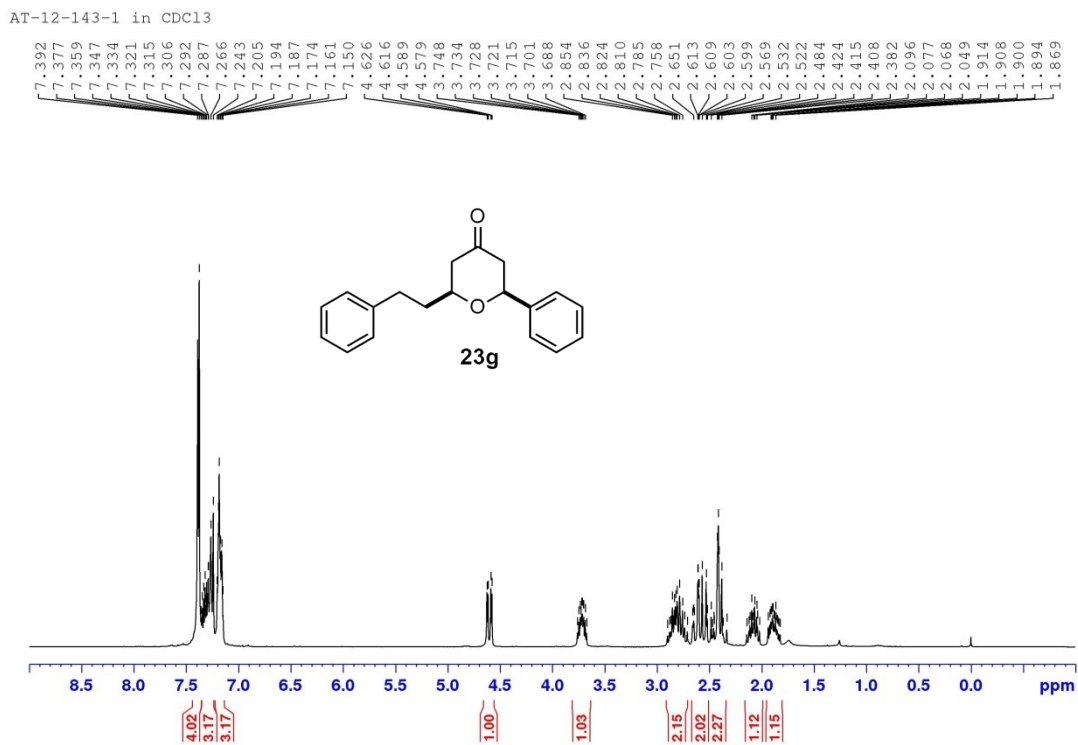


Figure SI22. ¹³C NMR (75 MHz, CDCl₃) spectrum of 2-phenethyl-6-phenyltetrahydropyran-4-one (**23g**)

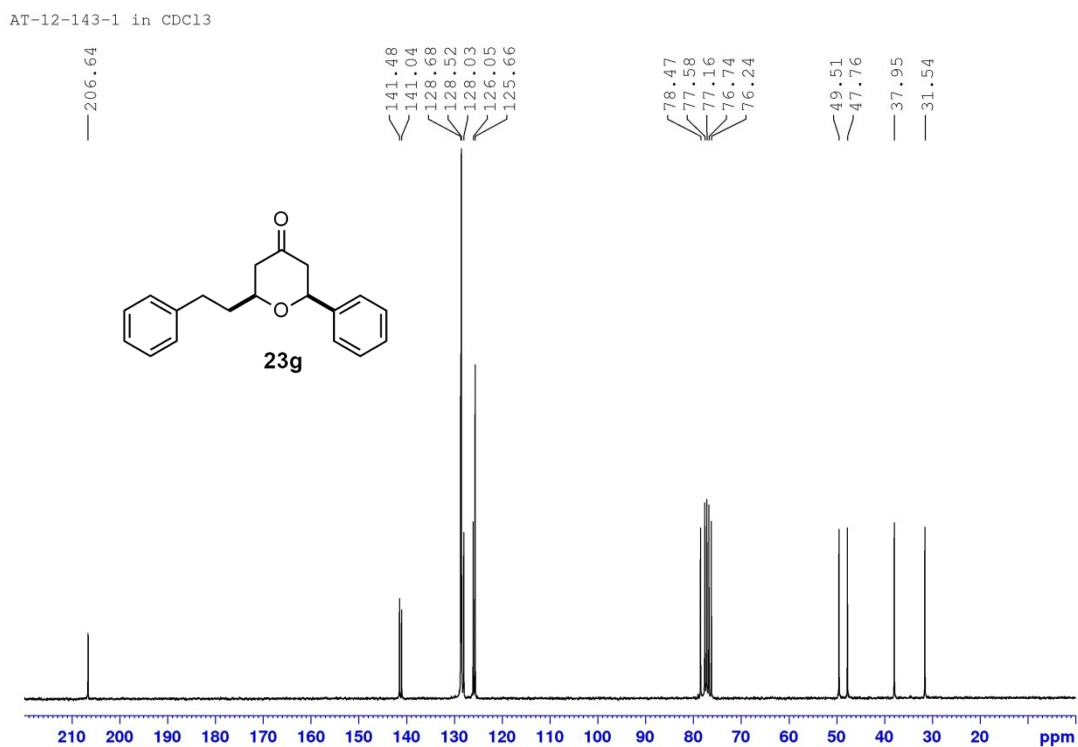


Figure SI23. ¹³C NMR (300 MHz, CDCl₃) spectrum of 2-(4-nitrophenyl)-6-phenethyl-tetrahydropyran-4-one (**23h**)

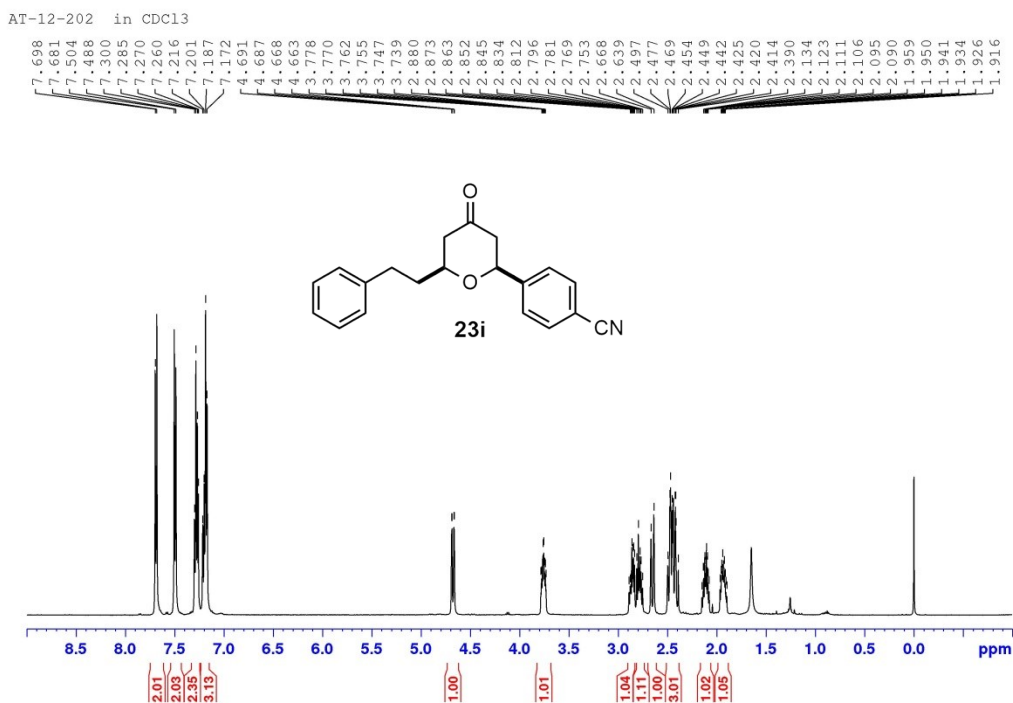


Figure SI26. ¹³C NMR (75 MHz, CDCl₃) spectrum of 2-(4-Cyanophenyl)-6-phenethyl-tetrahydropyran-4-one (**23i**)

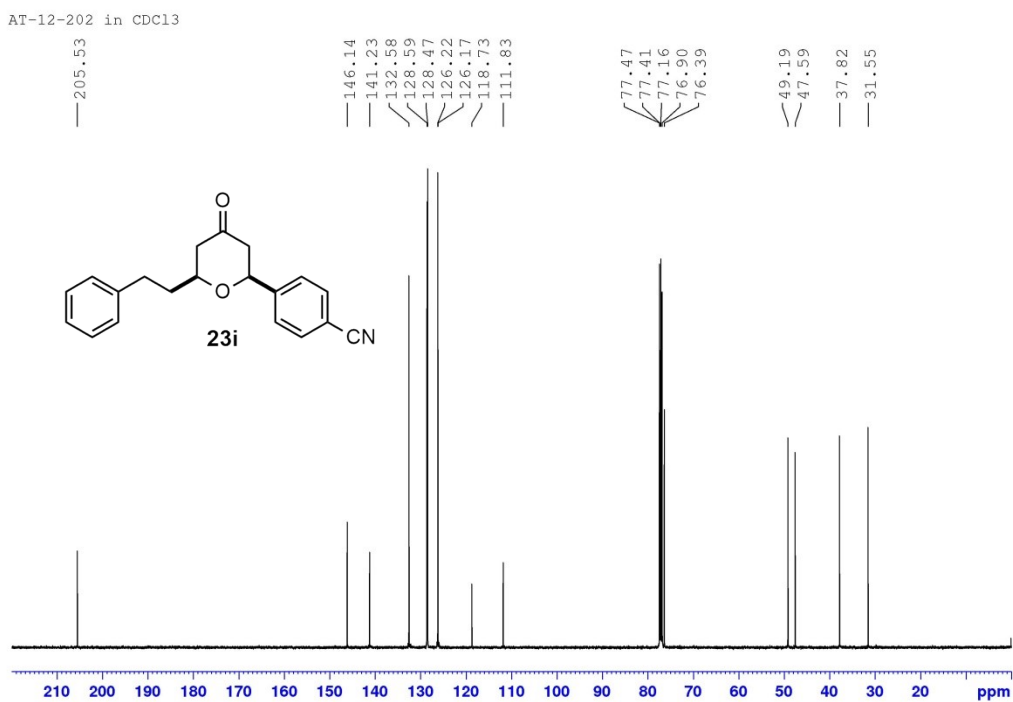


Figure SI27. ¹H NMR (300 MHz, CDCl₃) spectrum of 2-(4-bromophenyl)-6-phenethyl-tetrahydropyran-4-one (**23j**)

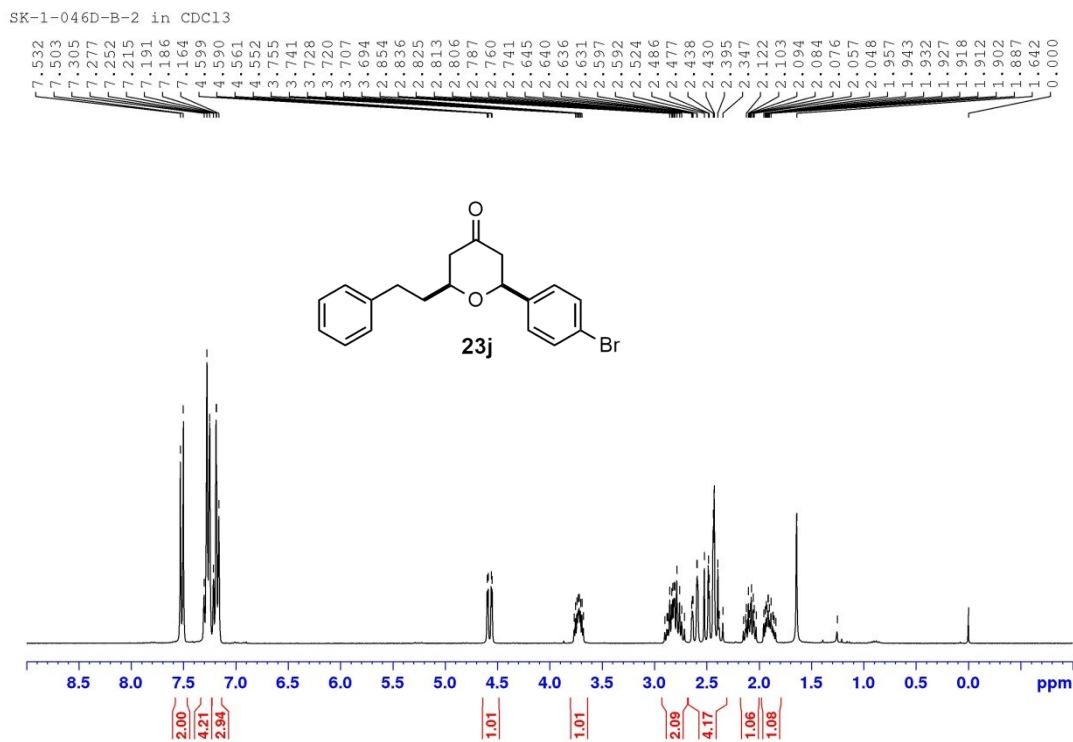


Figure SI28. ¹³C NMR (75 MHz, CDCl₃) spectrum of 2-(4-bromophenyl)-6-phenethyl-tetrahydropyran-4-one (**23j**)

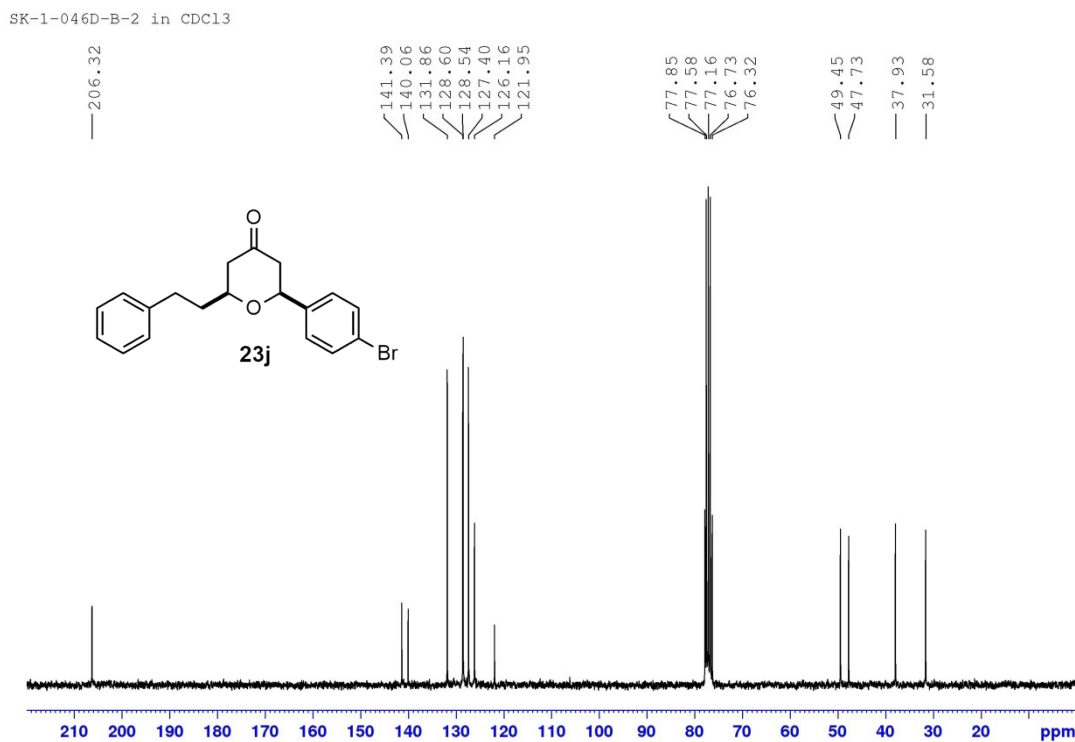


Figure SI29. ¹³C NMR (300 MHz, CDCl₃) spectrum of 2-(4-chlorophenyl)-6-phenethyl-tetrahydropyran-4-one (**23k**)

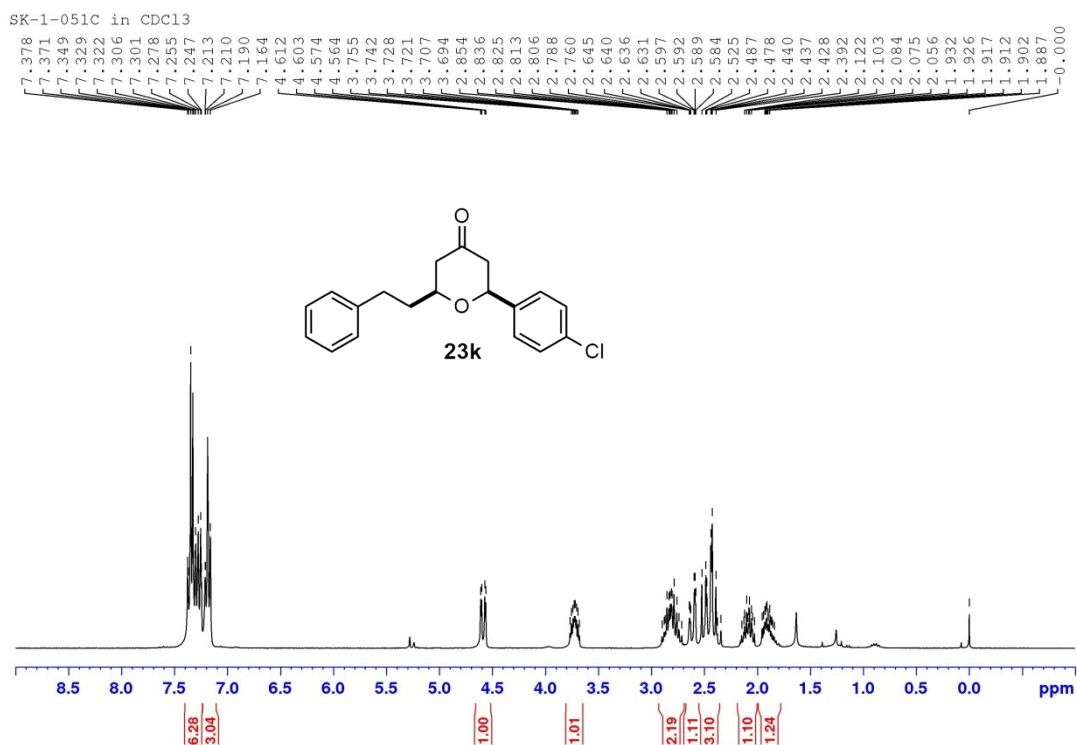


Figure SI30. ¹³C NMR (75 MHz, CDCl₃) spectrum of 2-(4-bromophenyl)-6-phenethyl-tetrahydropyran-4-one (**23k**)

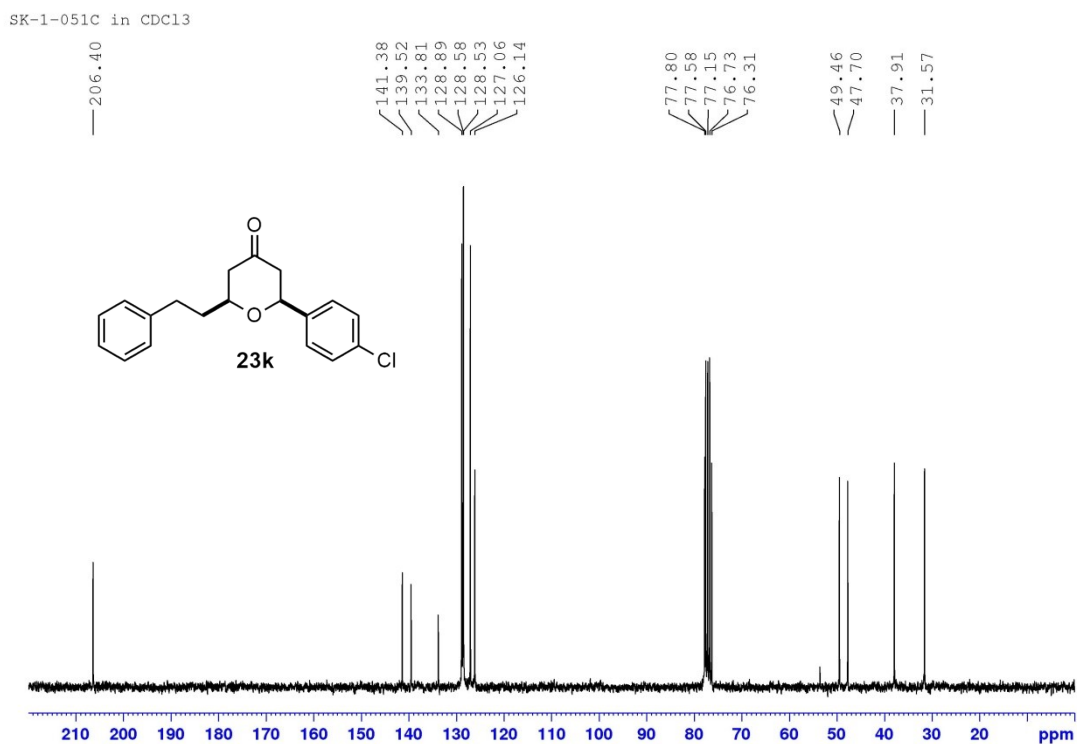


Figure SI31. ¹H NMR (300 MHz, CDCl₃) spectrum of 2-(4-methoxyphenyl)-6-phenethyltetrahydropyran-4-one (**23l**)

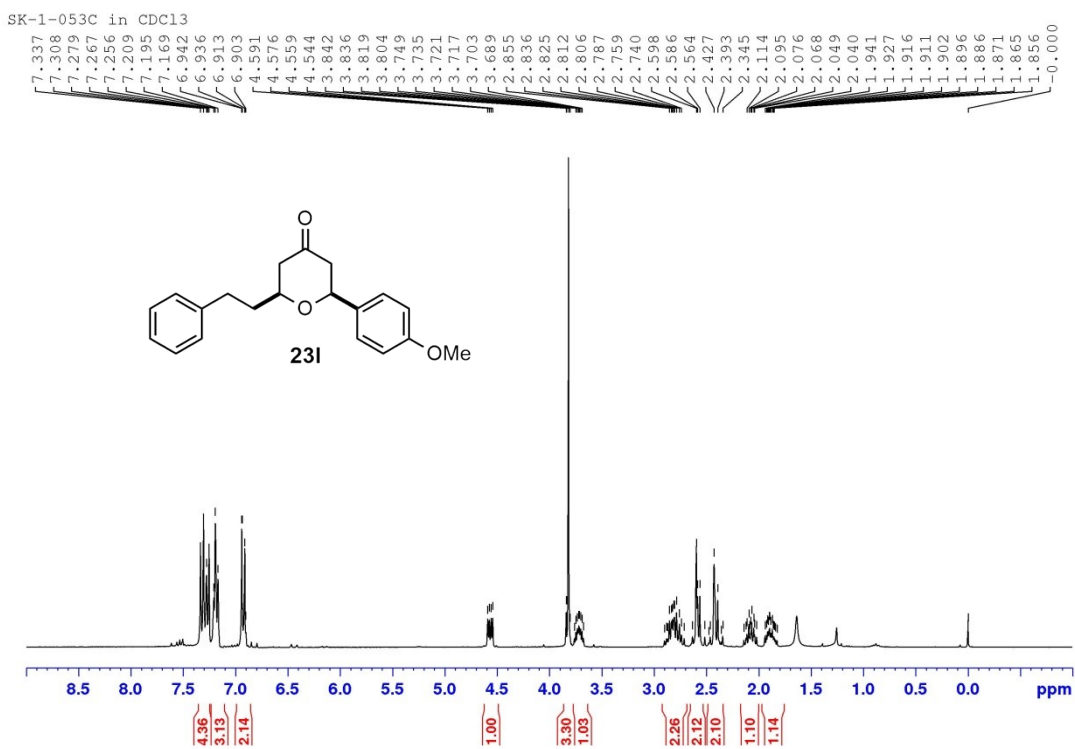


Figure SI32. ¹³C NMR (75 MHz, CDCl₃) spectrum of 2-(4-methoxyphenyl)-6-phenethyltetrahydropyran-4-one (**23l**)

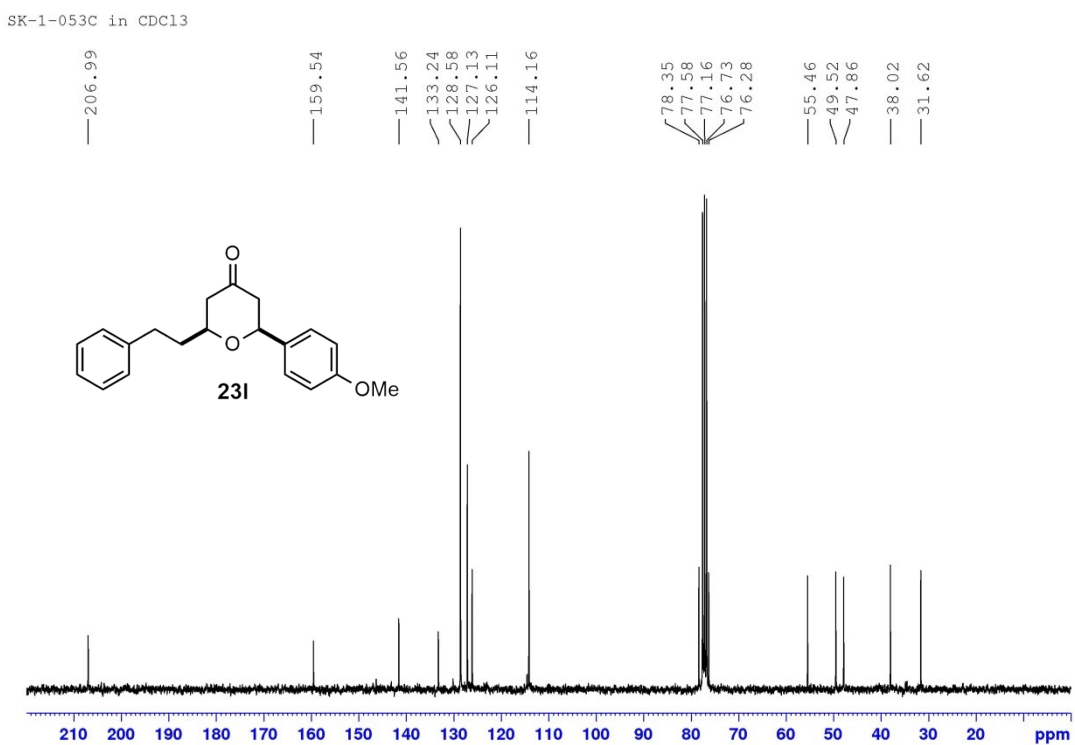


Figure SI33. ¹³C NMR (75 MHz, CDCl₃) spectrum of 2-(3-methoxyphenyl)-6-phenethyltetrahydropyran-4-one (**23m**)

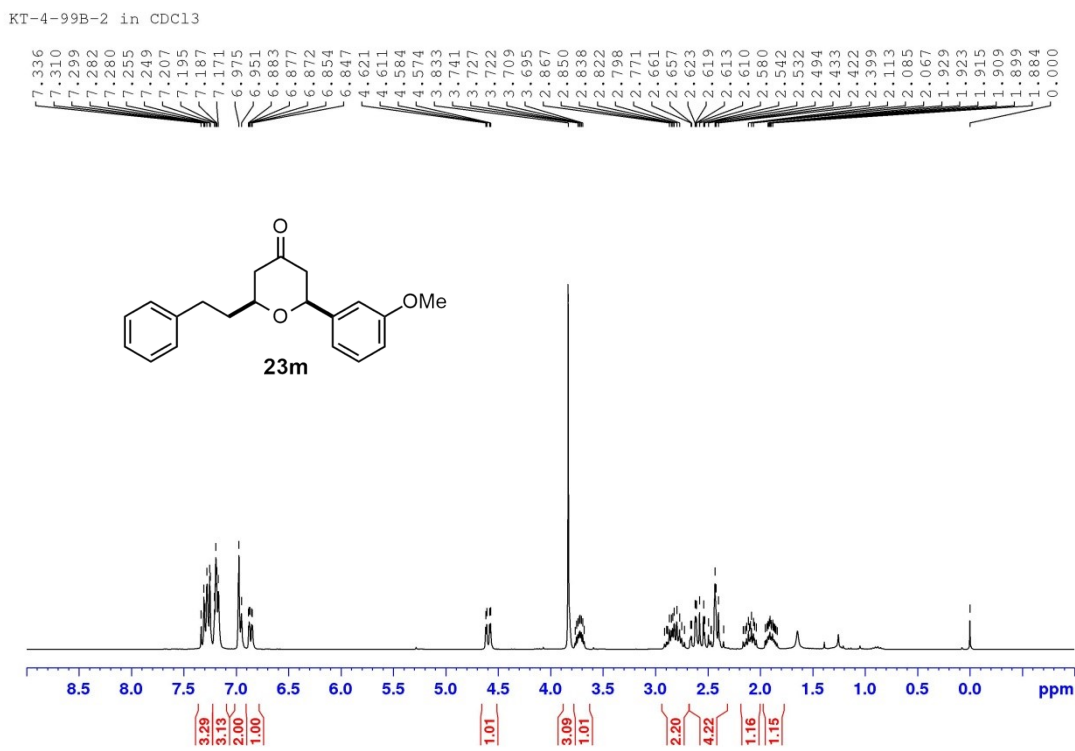


Figure SI34. ¹³C NMR (75 MHz, CDCl₃) spectrum of 2-(3-methoxyphenyl)-6-phenethyltetrahydropyran-4-one (**23m**)

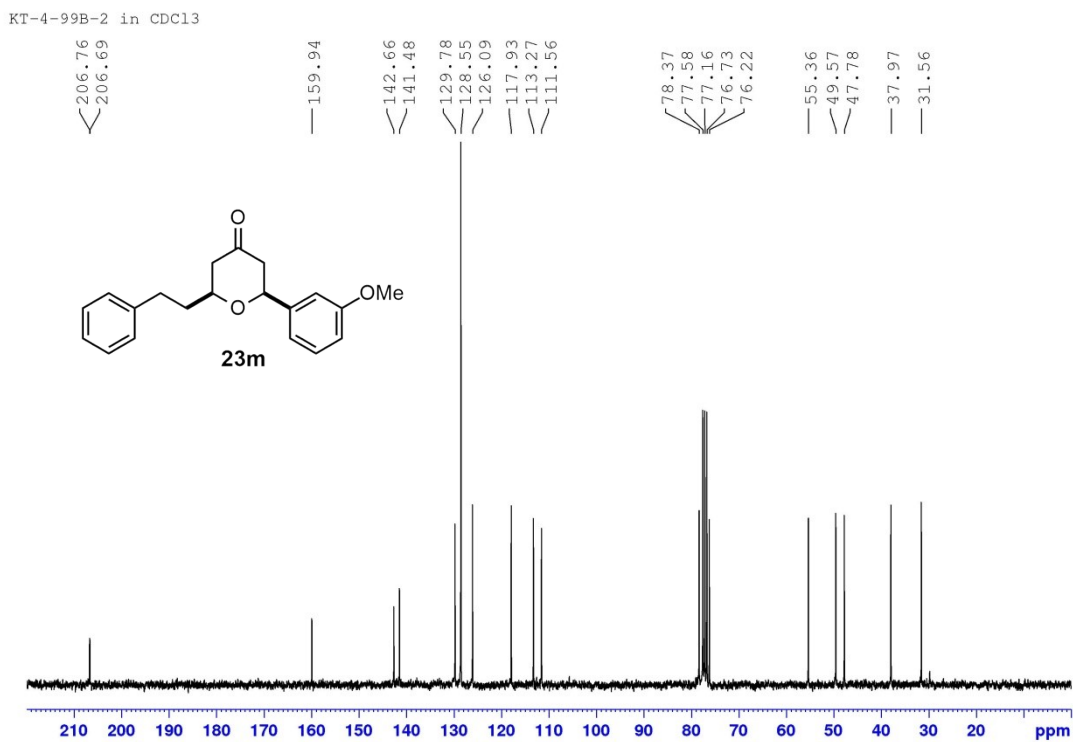


Figure SI35. ¹H NMR (300 MHz, CDCl₃) spectrum of 2-(2-methoxyphenyl)-6-phenethyltetrahydropyran-4-one (**23n**)

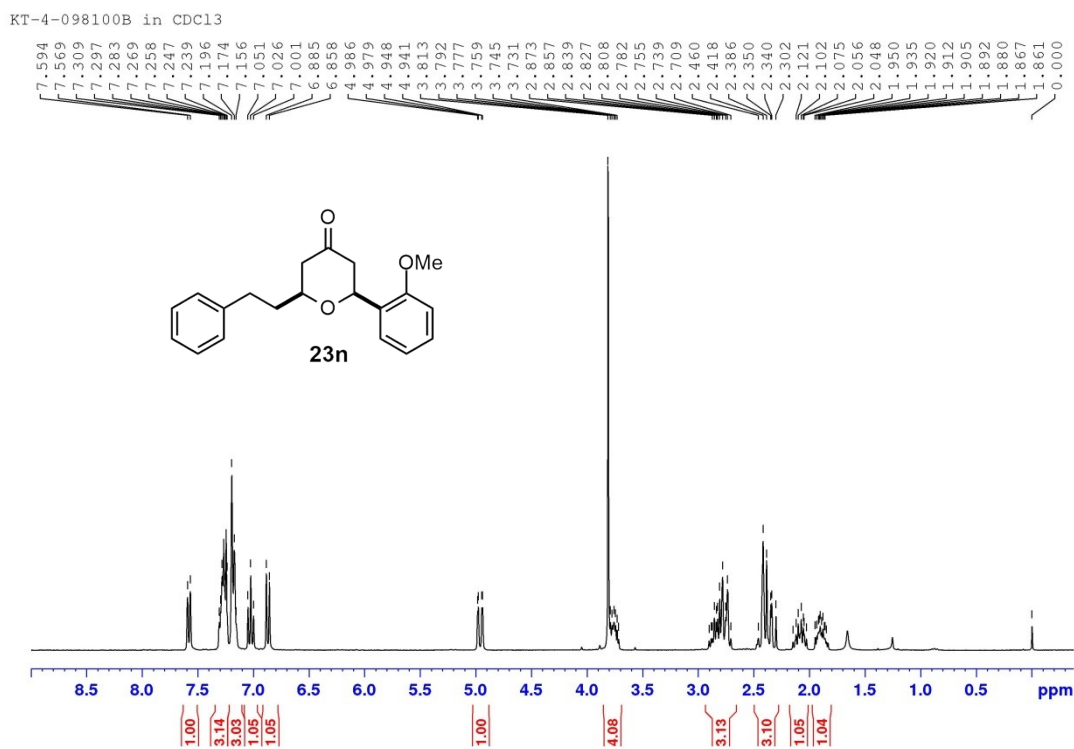


Figure SI36. ¹³C NMR (75 MHz, CDCl₃) spectrum of 2-(3-methoxyphenyl)-6-phenethyltetrahydropyran-4-one (**23n**)

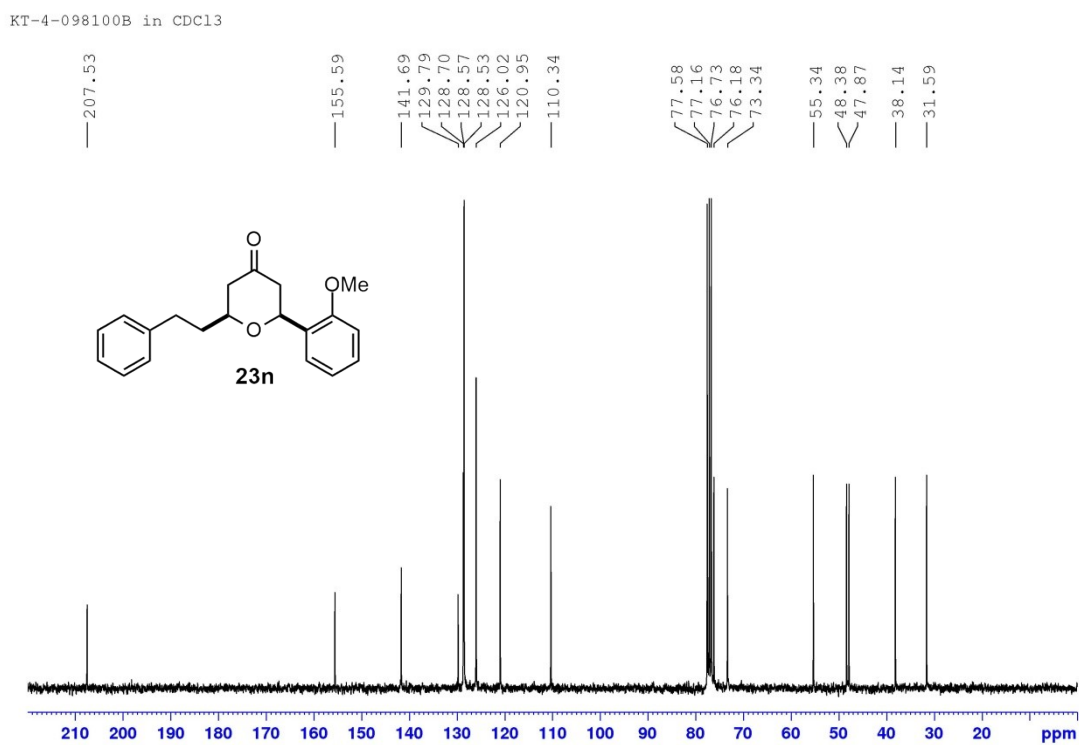


Figure SI37. ¹H NMR (300 MHz, CDCl₃) spectrum of 2-phenethyl-6-styryltetrahydropyran-4-one (**23o**)

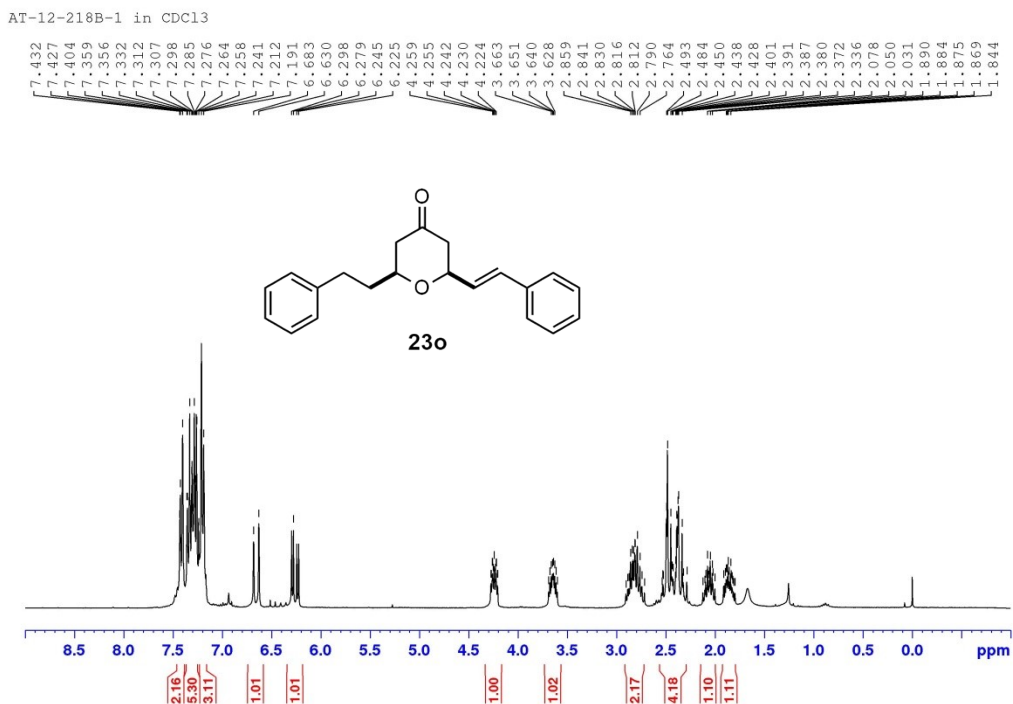


Figure SI38. ¹³C NMR (75 MHz, CDCl₃) spectrum of 2-phenethyl-6-styryltetrahydropyran-4-one (**23o**)

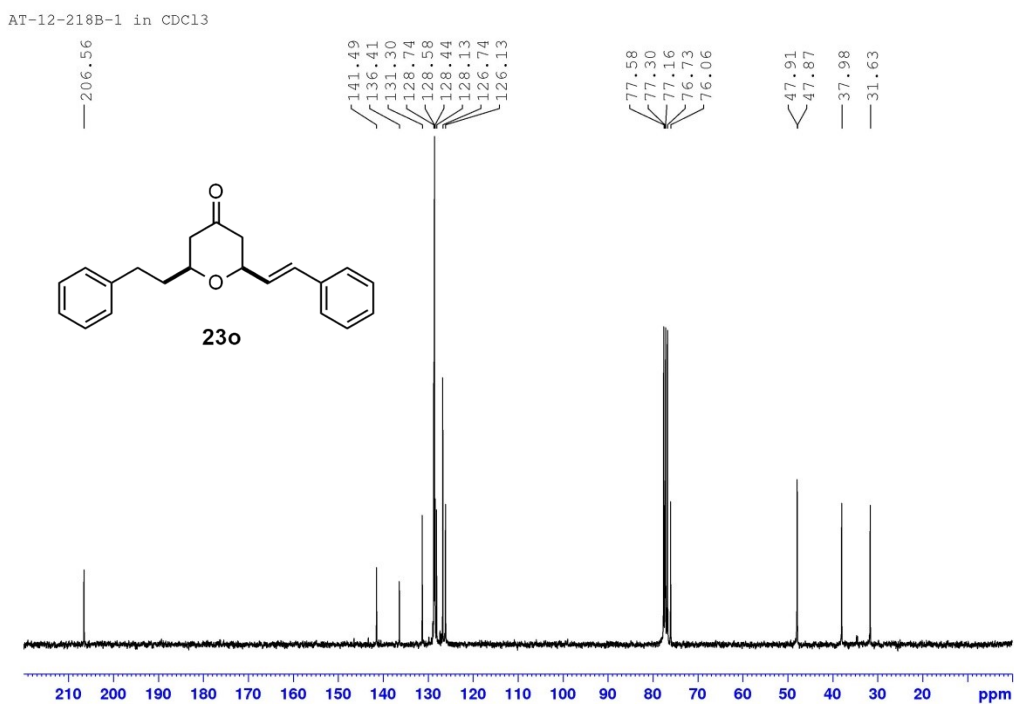


Figure SI39. ¹H NMR (300 MHz, CDCl₃) spectrum of 2-(3-benzoyloxypropyl)-6-cyclohexyltetrahydropyran-4-one (**23p**)

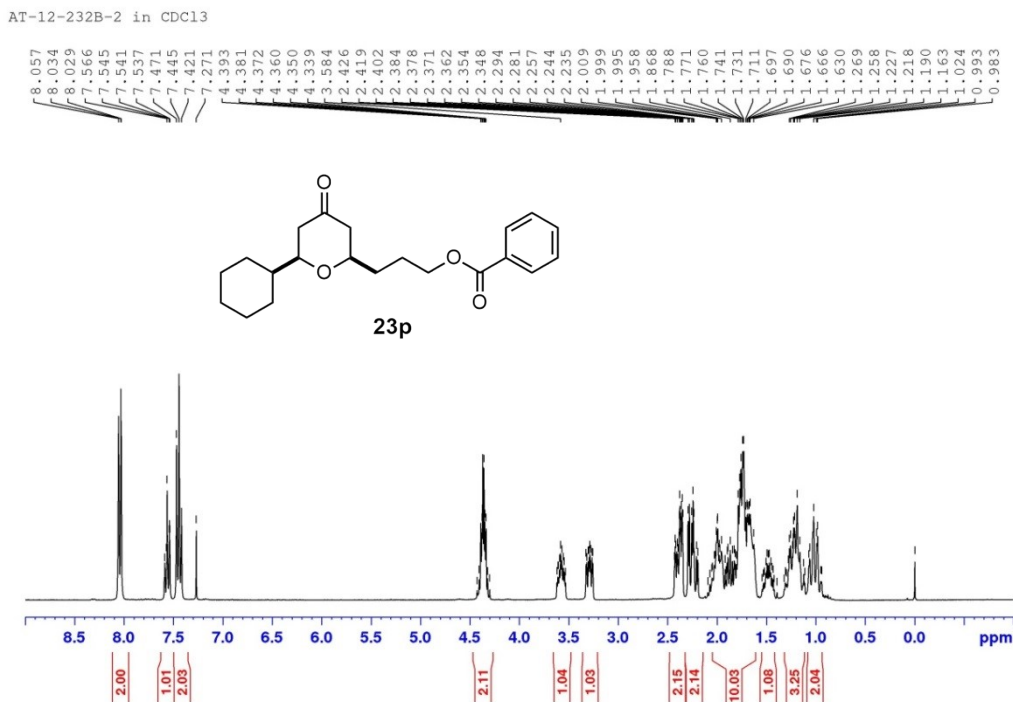


Figure SI40. ¹³C NMR (75 MHz, CDCl₃) spectrum of 2-(3-benzoyloxypropyl)-6-cyclohexyltetrahydropyran-4-one (**23p**)

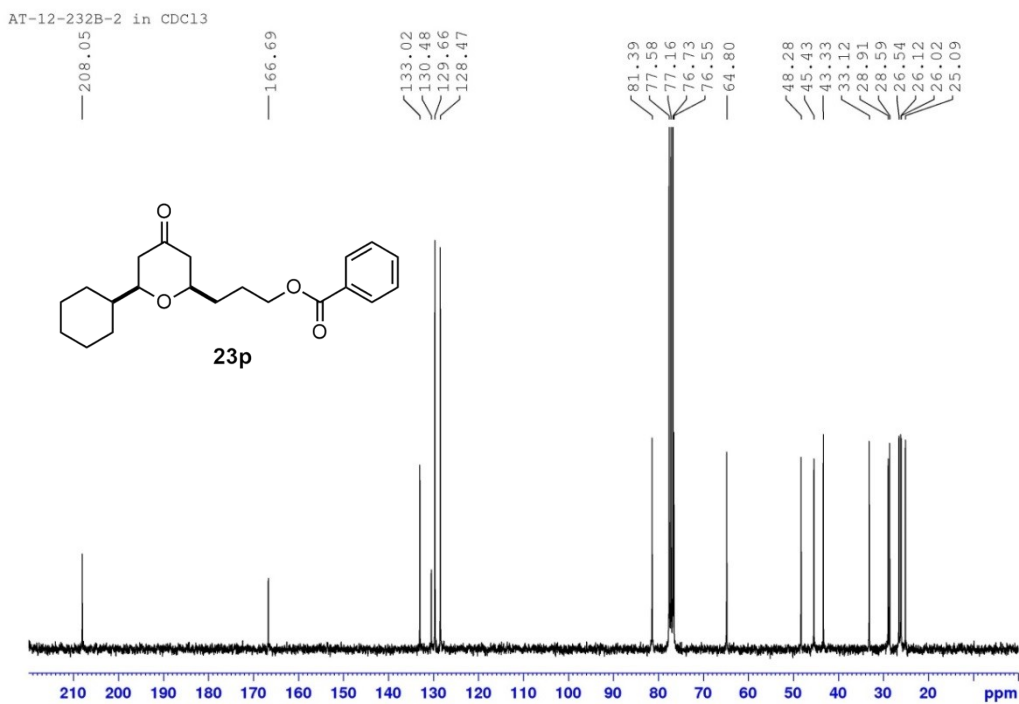


Figure SI41. ¹H NMR (300 MHz, CDCl₃) spectrum of 2-cyclohexyl-6-propyltetrahydropyran-4-one (**23q**)

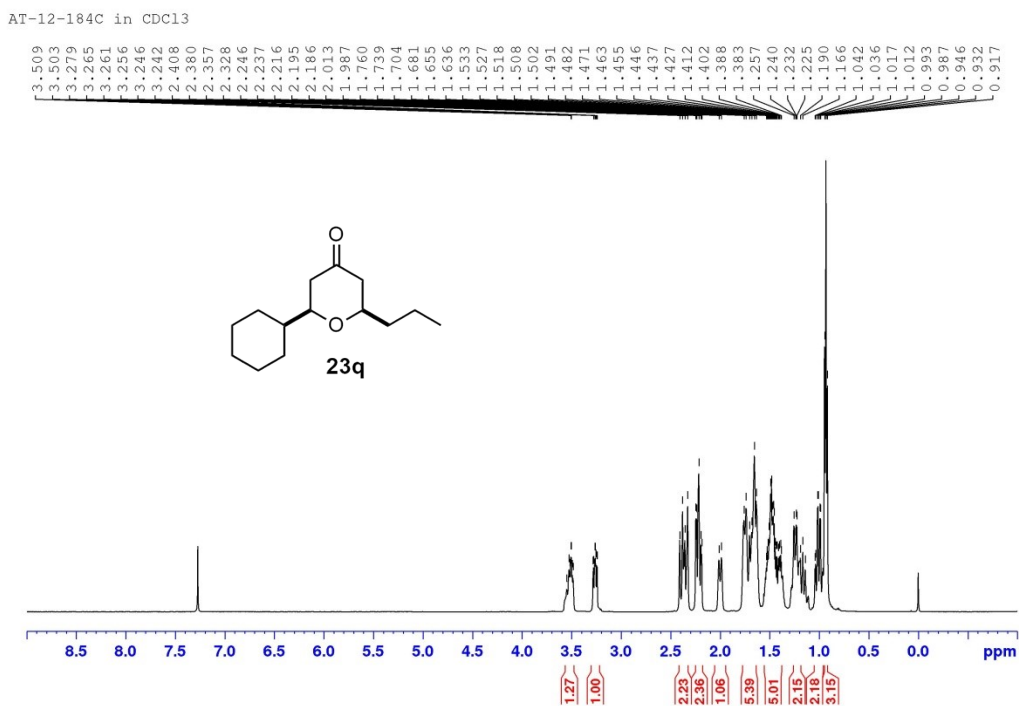


Figure SI42. ¹³C NMR (75 MHz, CDCl₃) spectrum of 2-cyclohexyl-6-propyltetrahydropyran-4-one (**23q**)

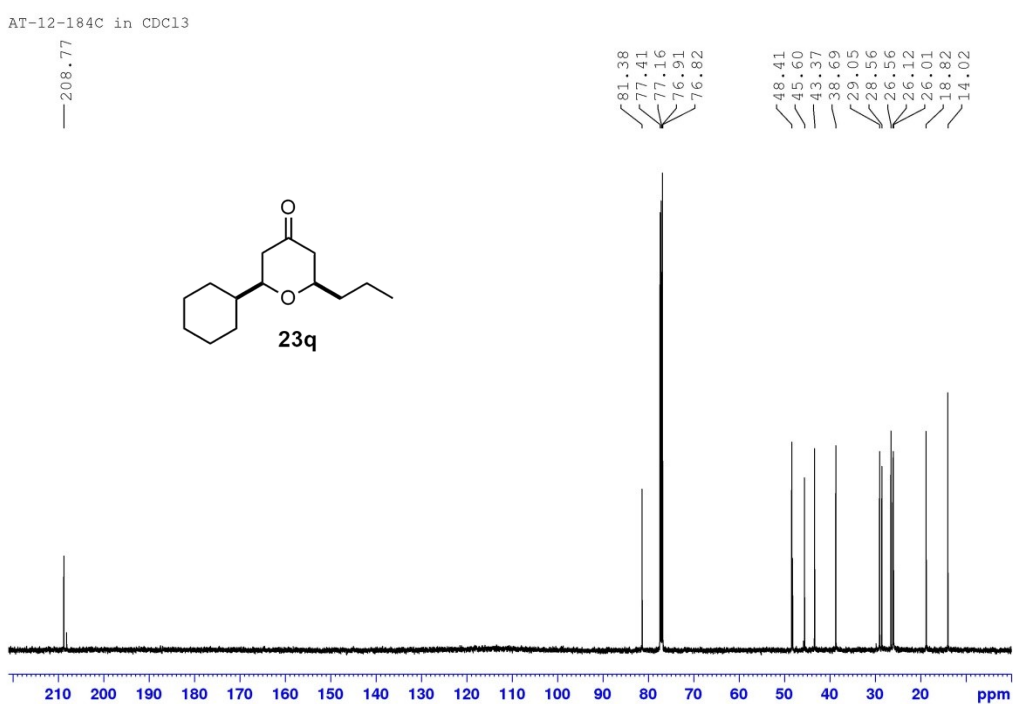


Figure SI43. ¹H NMR (300 MHz, CDCl₃) spectrum of 2,6-dicyclohexyltetrahydropyran-4-one (**23r**)

SK-1-071B in CDCl₃

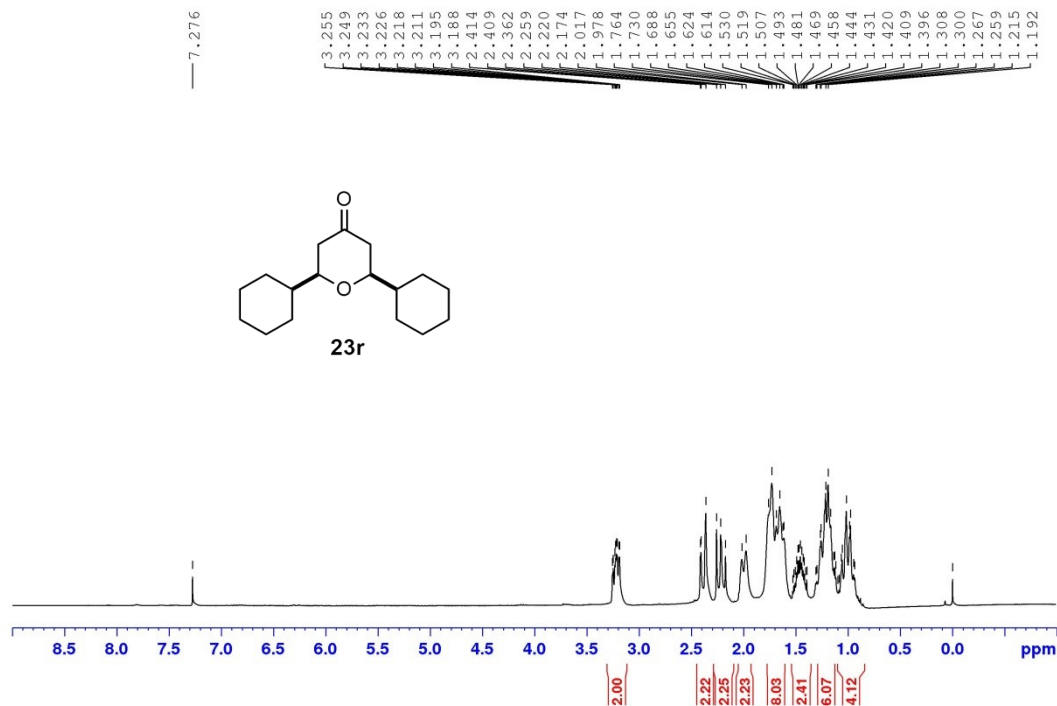


Figure SI44. ¹³C NMR (75 MHz, CDCl₃) spectrum of 2,6-dicyclohexyltetrahydropyran-4-one (**23r**)

SK-1-071B in CDCl₃

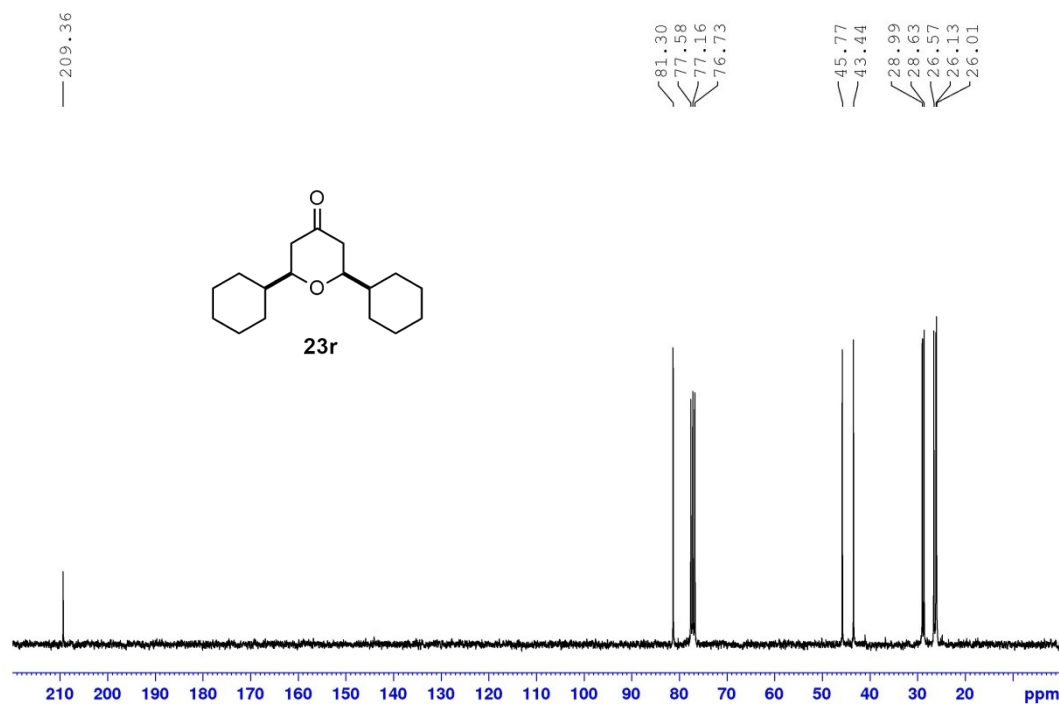


Figure SI45. ¹H NMR (500 MHz, CDCl₃) spectrum of 2-cyclohexyl-6-(4-nitrophenyl) tetrahydropyran-4-one (**23s**)

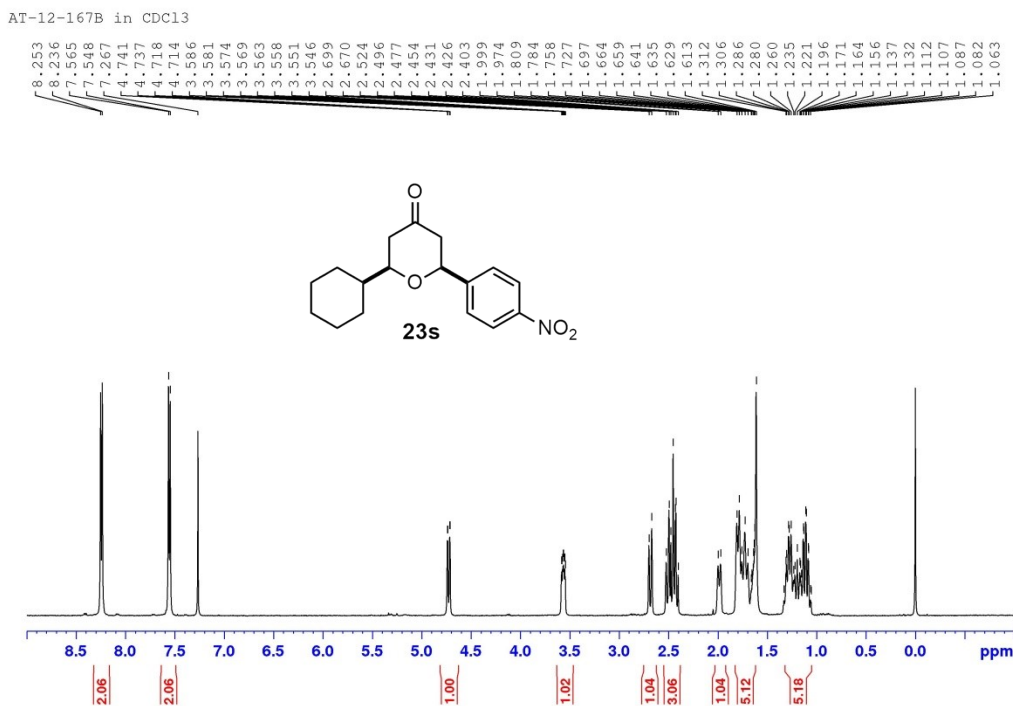


Figure SI46. ¹³C NMR (125 MHz, CDCl₃) spectrum of 2-cyclohexyl-6-(4-nitrophenyl) tetrahydropyran-4-one (**23s**)

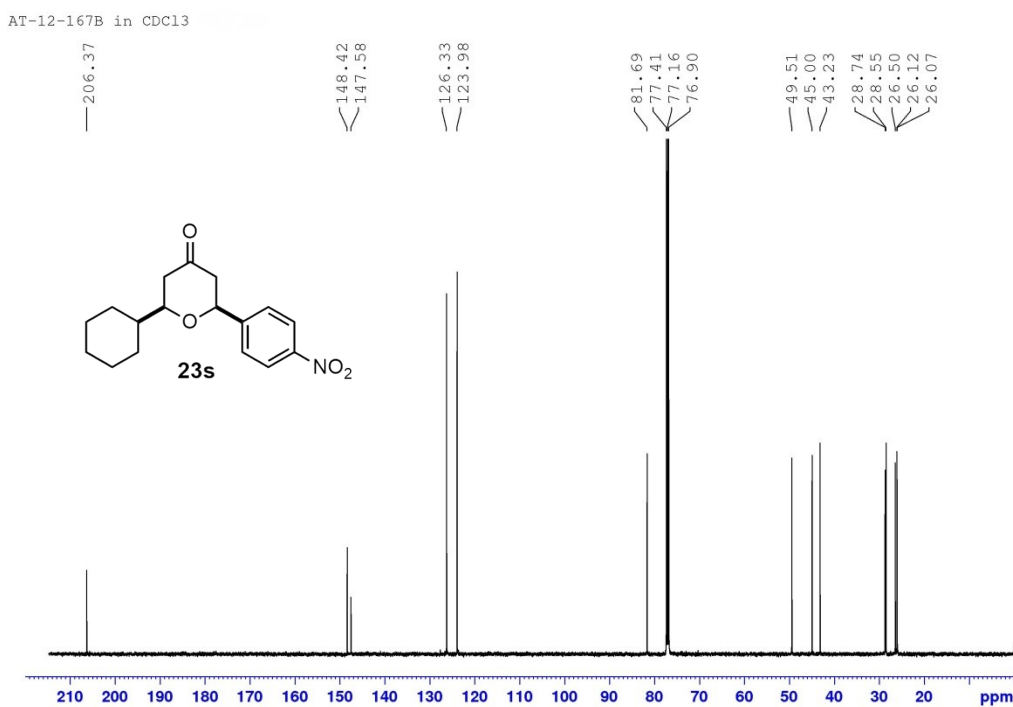


Figure SI47. ¹H NMR (300 MHz, CDCl₃) spectrum of 2-Cyclohexyl-6-(4-methoxyphenyl)tetrahydropyran-4-one (**23t**)

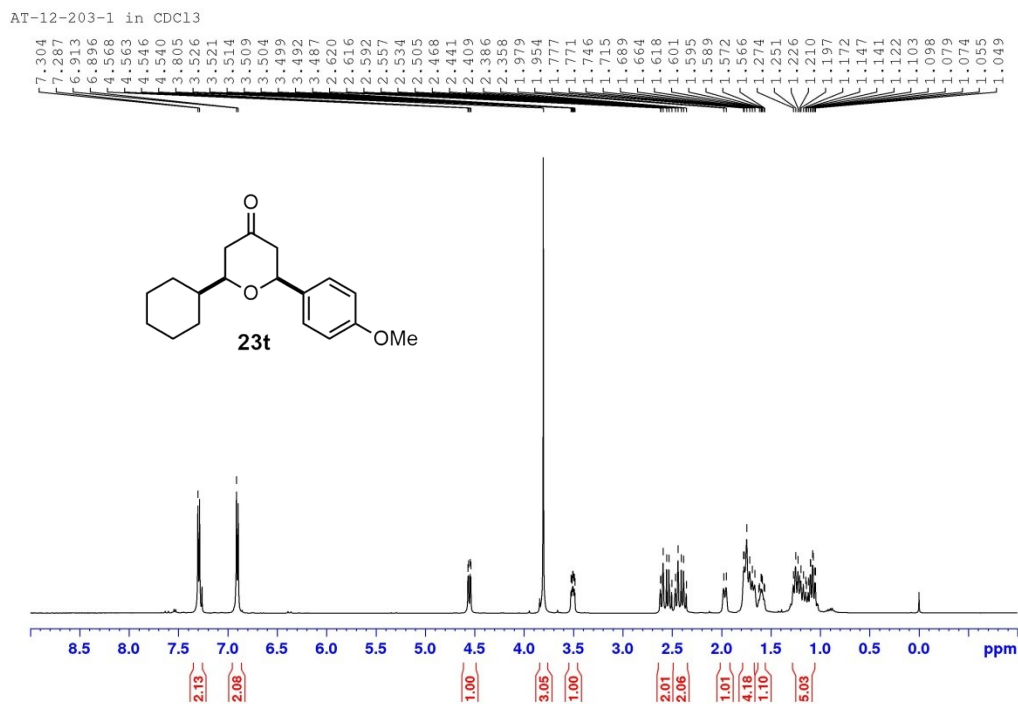


Figure SI48. ¹³C NMR (75 MHz, CDCl₃) spectrum of 2-cyclohexyl-6-(4-methoxyphenyl)tetrahydropyran-4-one (**23t**)

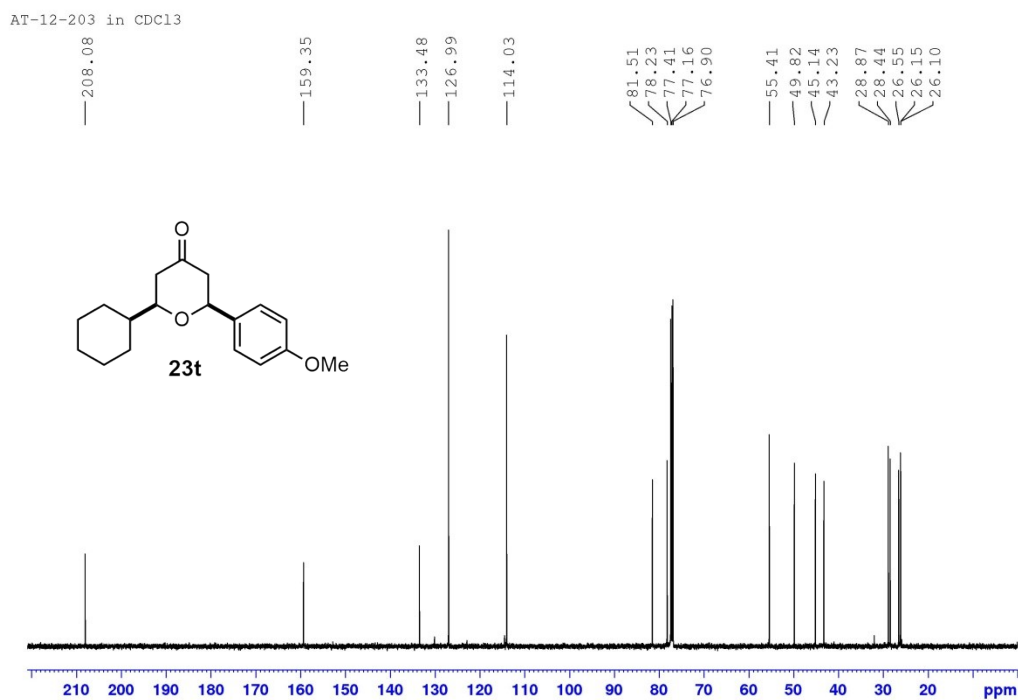


Figure SI49. ¹H NMR (300 MHz, CDCl₃) spectrum of 2-isopropyl-6-phenyltetrahydropyran-4-one (**23u**)

PV-3-036C in CDCl₃

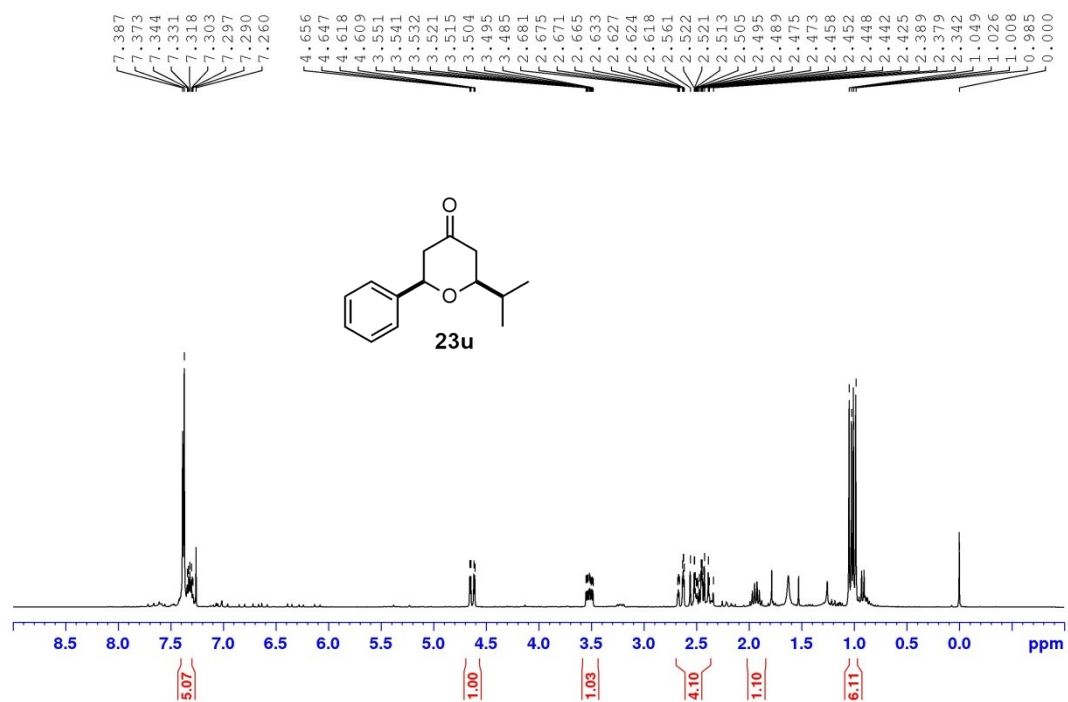


Figure SI50. ¹³C NMR (75 MHz, CDCl₃) spectrum of 2-isopropyl-6-phenyltetrahydropyran-4-one (**23u**)

AT-12-213A-2 in CDCl₃

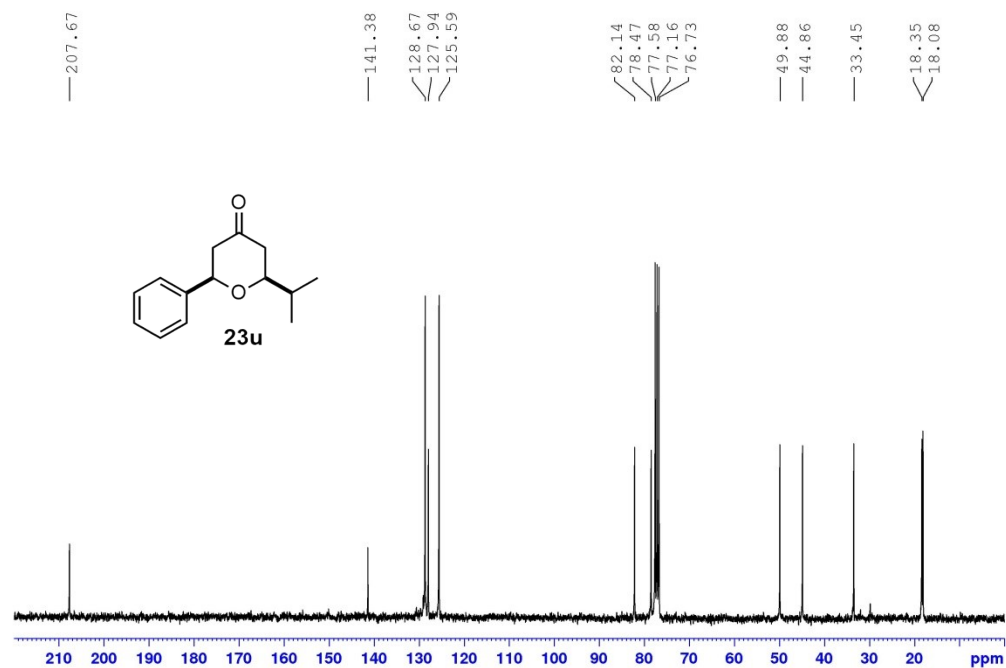


Figure SI51. ¹H NMR (300 MHz, CDCl₃) spectrum of 2,6-diisopropyltetrahydropyran-4-one (**23v**)

AT-12-220A in CDCl₃

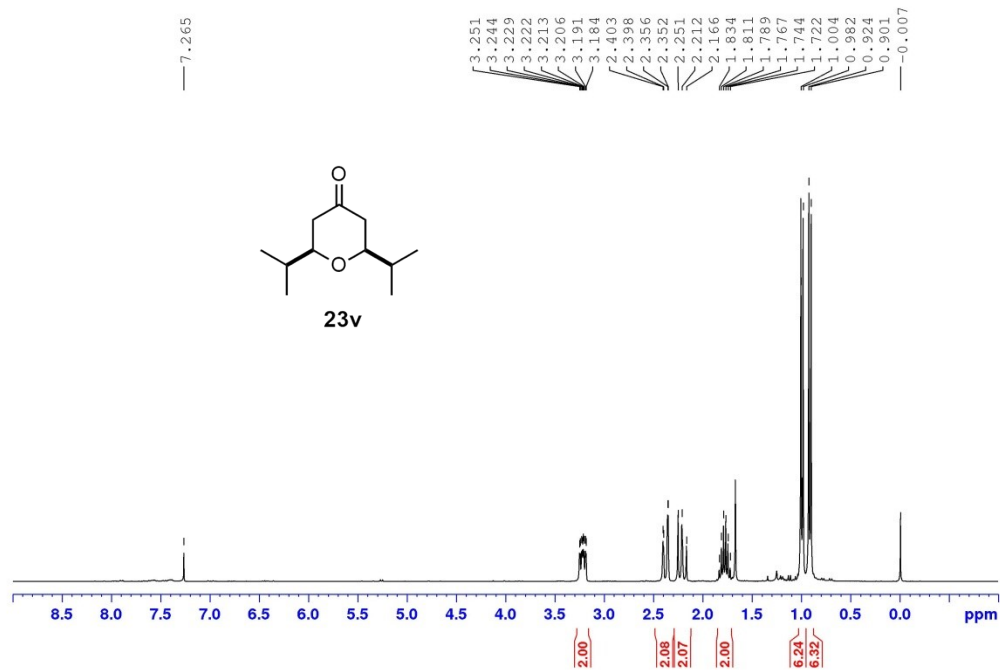


Figure SI52. ¹³C NMR (75 MHz, CDCl₃) spectrum of 2,6-diisopropyltetrahydropyran-4-one (**23v**)

AT-12-220A in CDCl₃

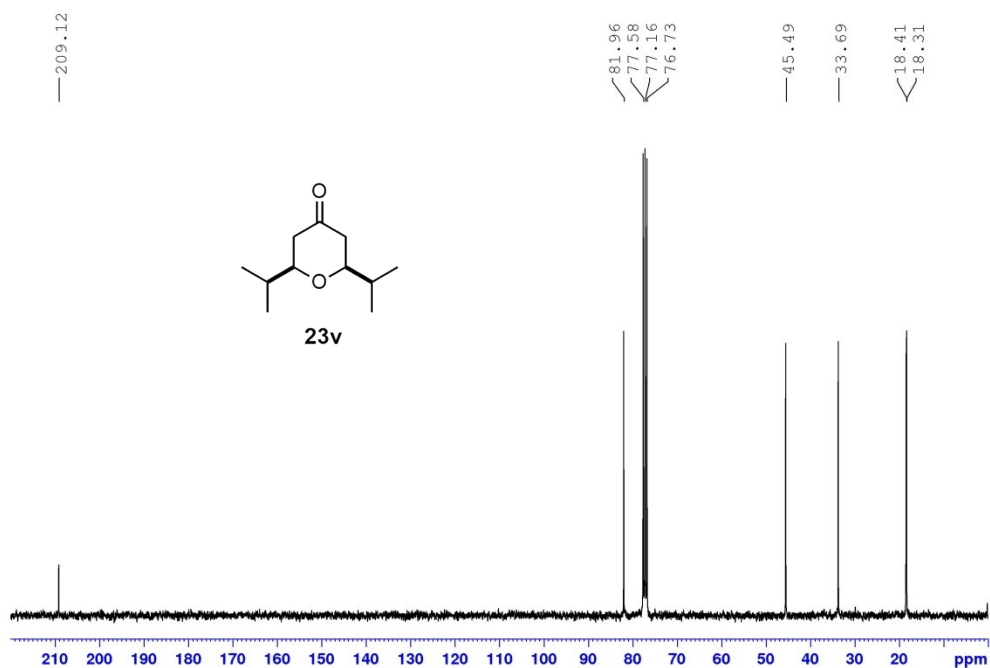


Figure SI53. ¹H NMR (300 MHz, CDCl₃) spectrum of 2,6-diphenyltetrahydropyran-4-one (**23w**)

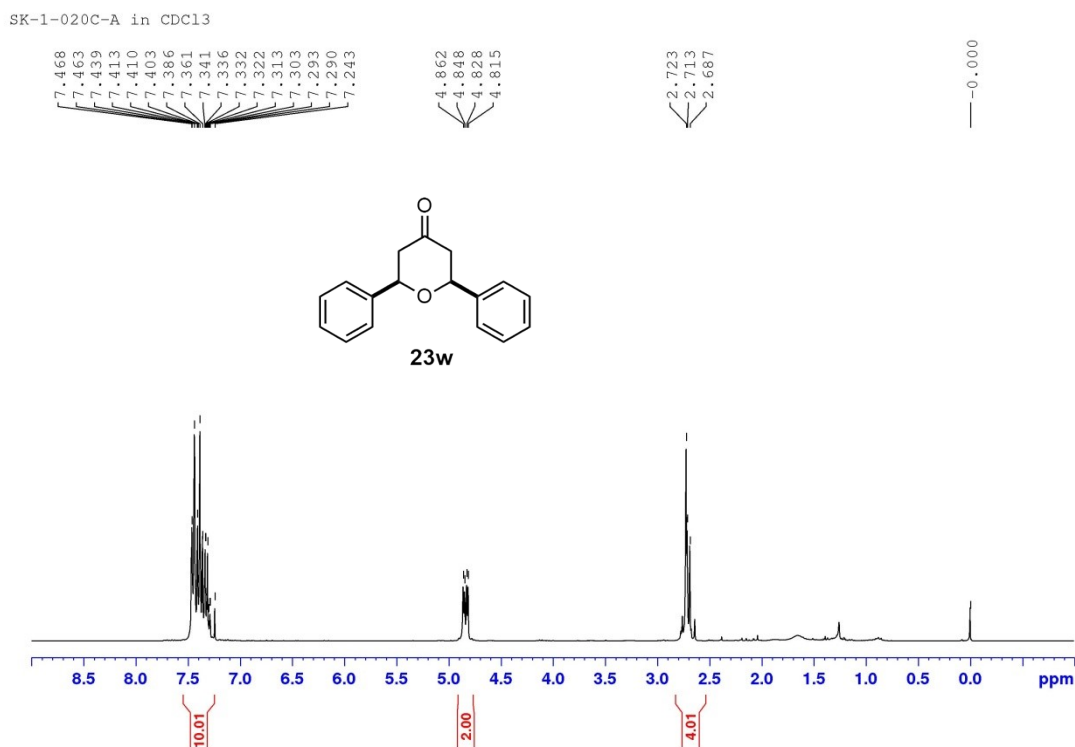


Figure SI54. ¹³C NMR (75 MHz, CDCl₃) spectrum of 2,6-diphenyltetrahydropyran-4-one (**23w**)

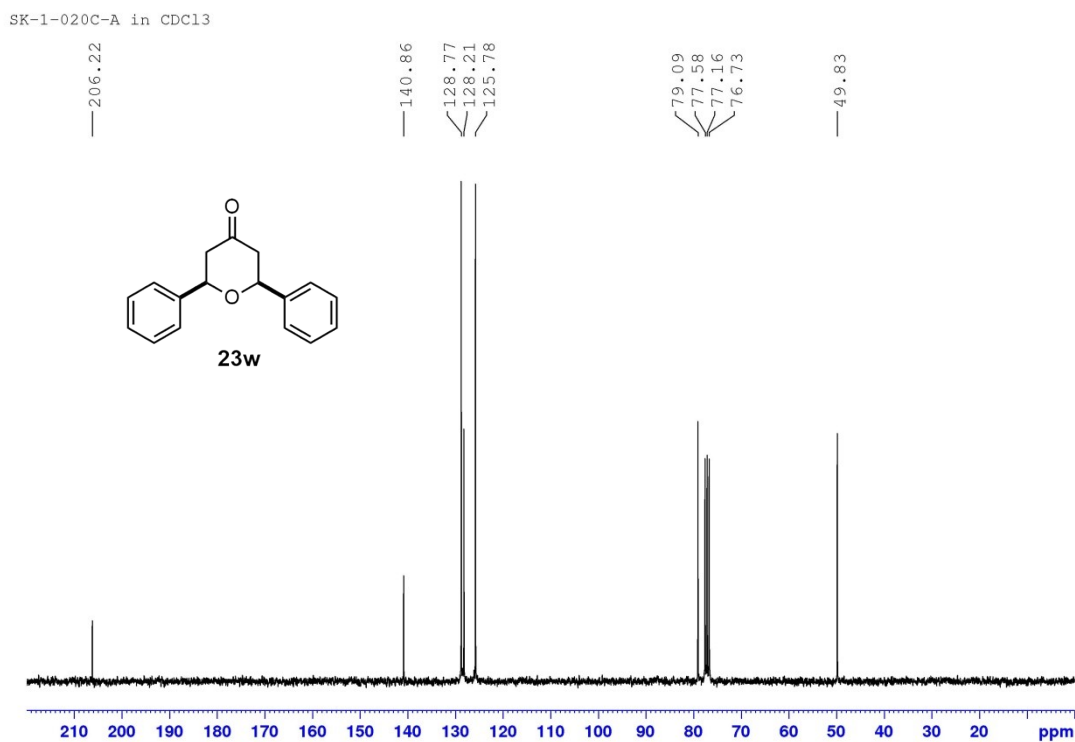


Figure SI55. ¹H NMR (300 MHz, CDCl₃) spectrum of 2-(4-nitrophenyl)-6-phenyltetrahydropyran-4-one (**23x**)

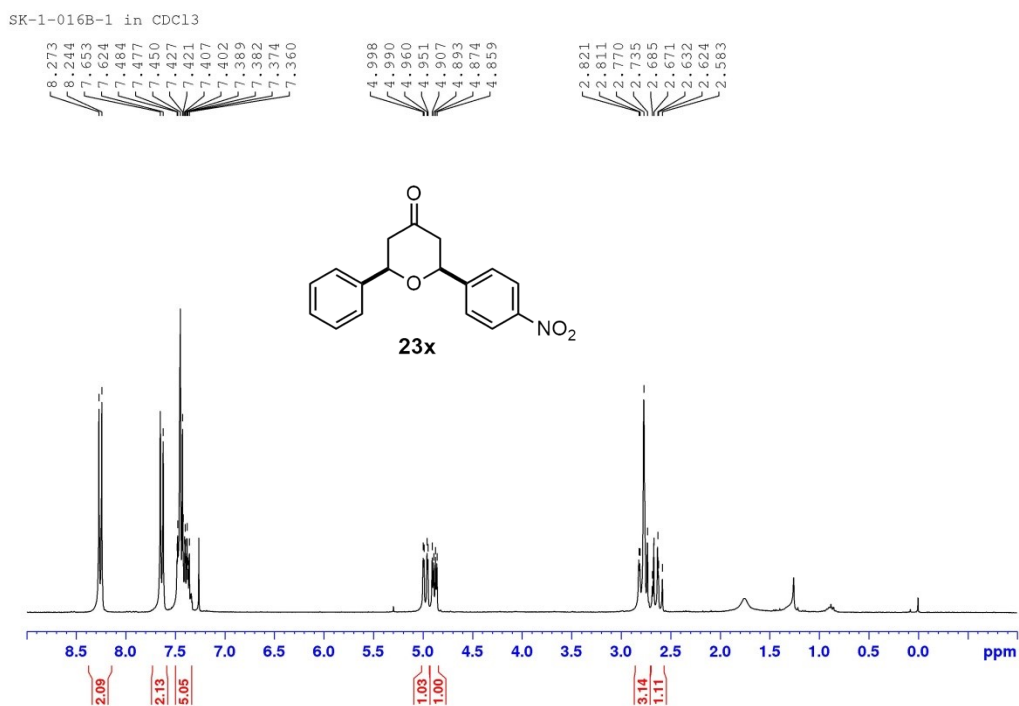


Figure SI56. ¹³C NMR (75 MHz, CDCl₃) spectrum of 2-(4-nitrophenyl)-6-phenyltetrahydropyran-4-one (**23x**)

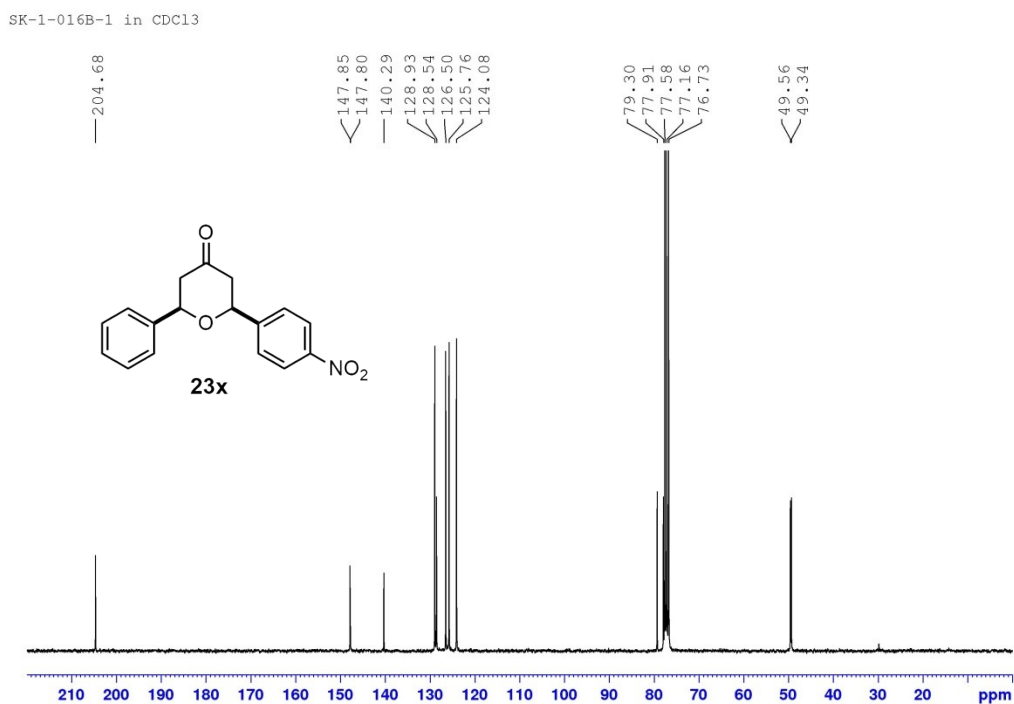


Figure SI57. ¹H NMR (300 MHz, CDCl₃) spectrum of 2-(3-methoxyphenyl)-6-phenyltetrahydropyran-4-one (**23y**)

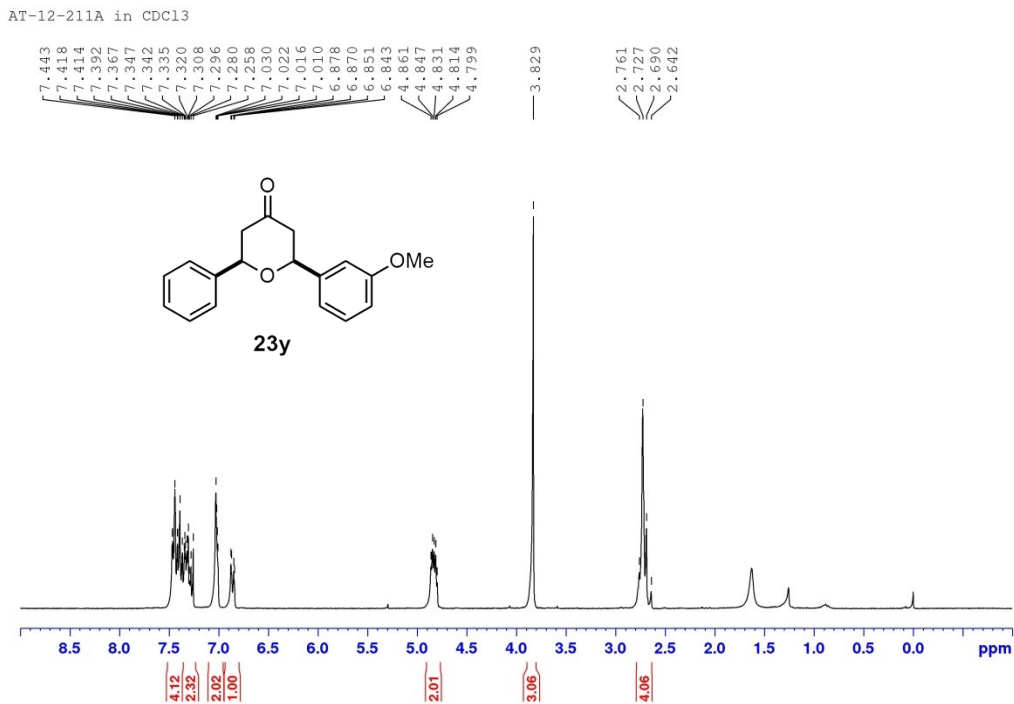


Figure SI58. ¹³C NMR (75 MHz, CDCl₃) spectrum of 2-(3-methoxyphenyl)-6-phenyl-tetrahydropyran-4-one (**23y**)

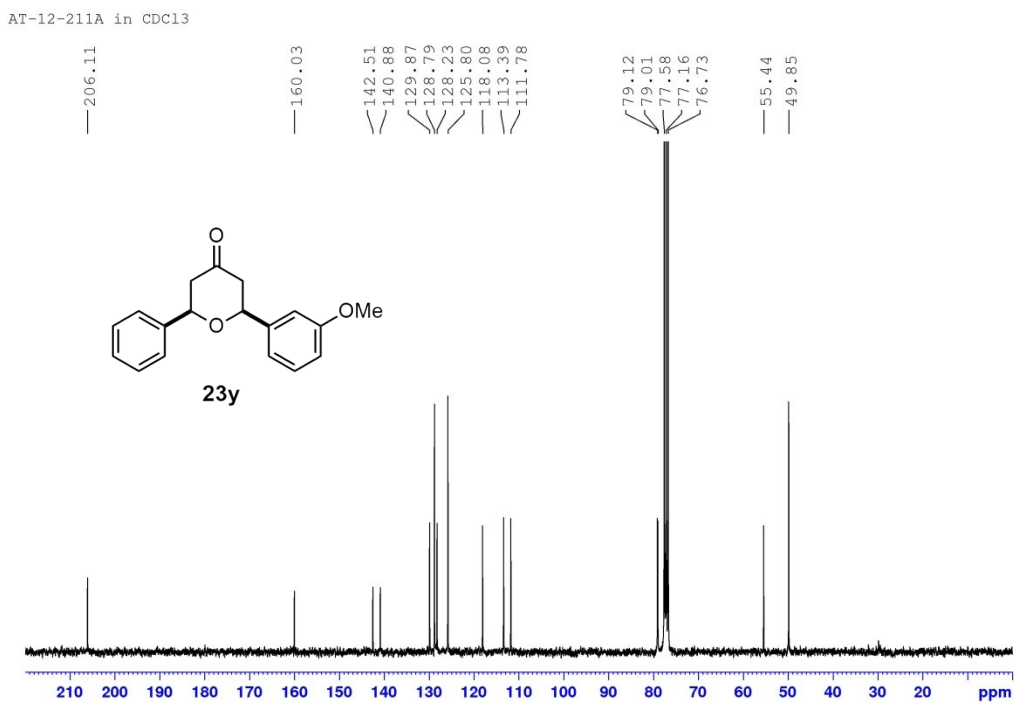


Figure SI59. ¹H NMR (300 MHz, CDCl₃) spectrum of 3-chloro-1-(3-methoxyphenyl)but-3-en-1-ol (**23z**)

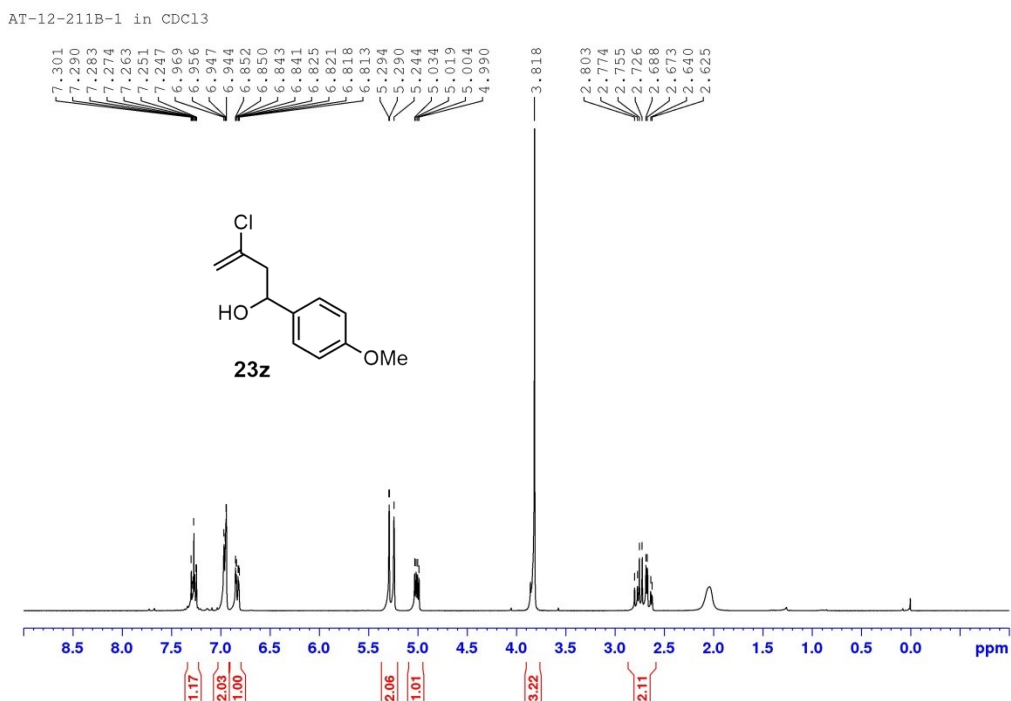
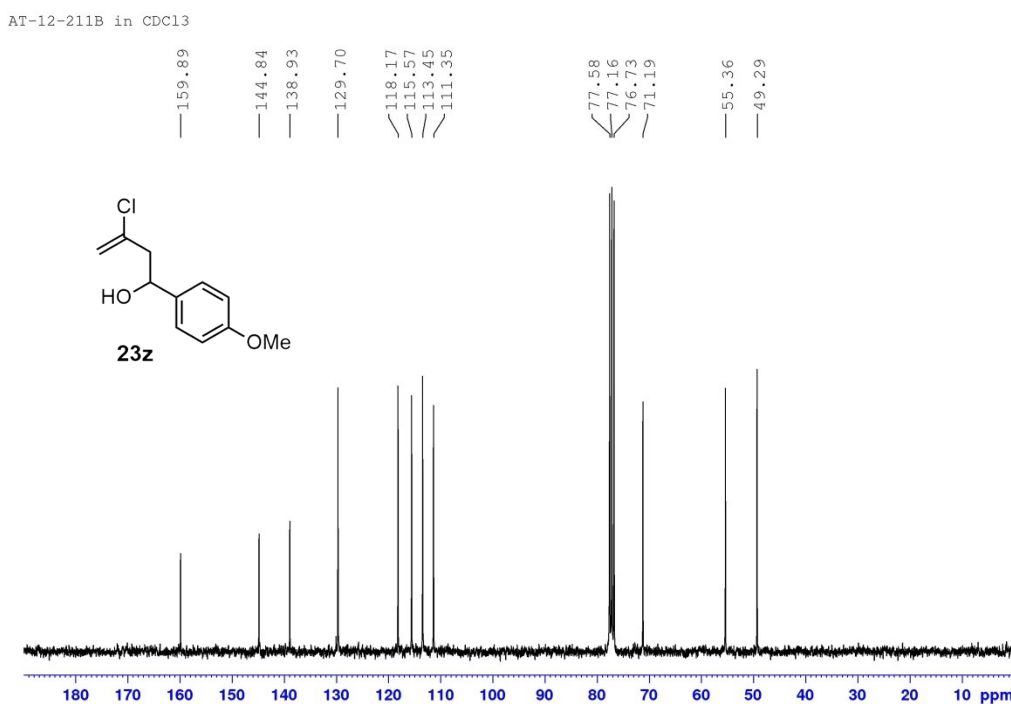


Figure SI60. ¹³C NMR (75 MHz, CDCl₃) spectrum of 3-chloro-1-(3-methoxyphenyl)but-3-en-1-ol (**23z**)



4. NOE Spectra of Selected Compounds

Figure SI61. NOE spectra (A) irradiation of ^1H at 3.23 ppm B) irradiation of ^1H at 3.49 ppm of 2-isopropyl-6-phenethyltetrahydropyran-4-one (**23e**)

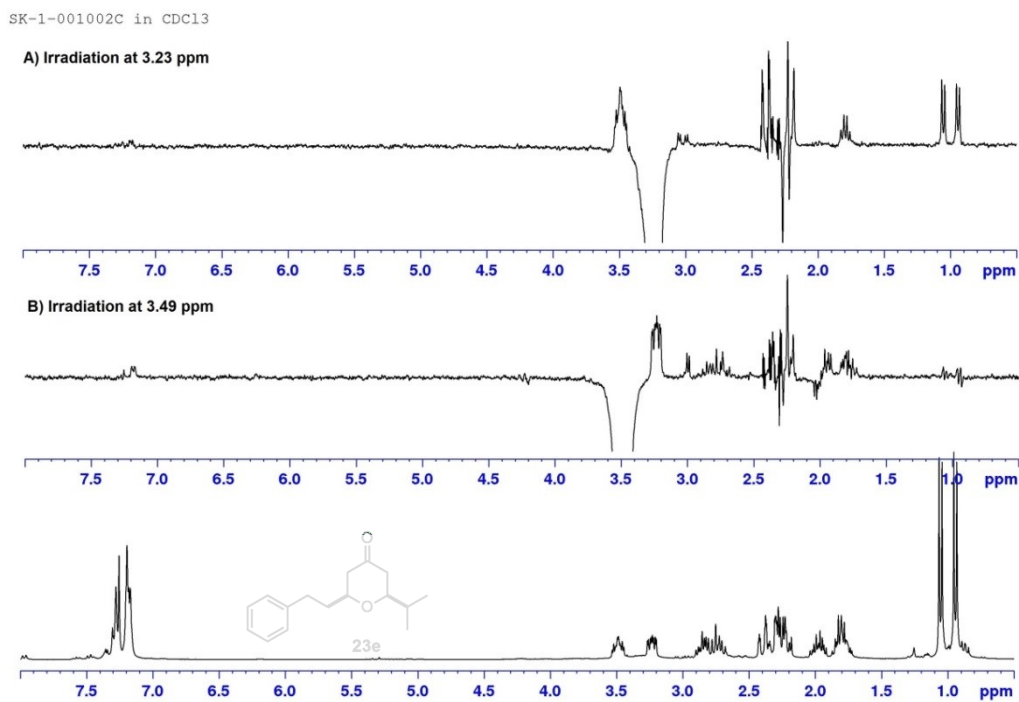
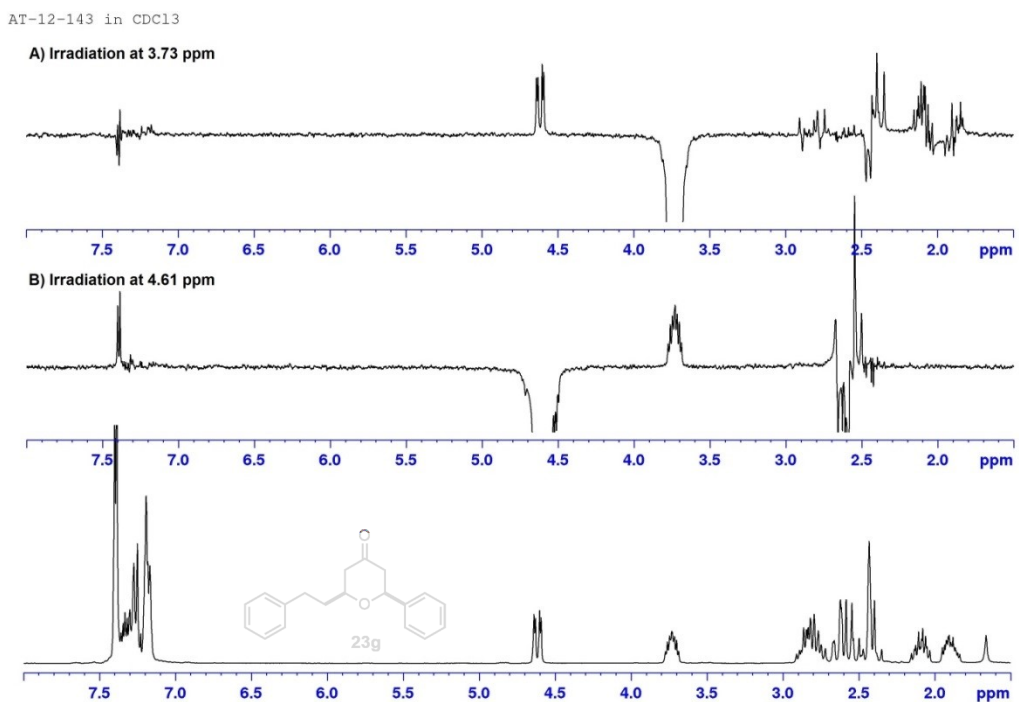


Figure SI62. NOE spectra (A) irradiation of ^1H at 3.73 ppm B) irradiation of ^1H at 4.61 ppm of 2-phenethyl-6-phenyltetrahydropyran-4-one (**23g**)



5. References

1. X.-H. Yi, Y. Meng and C.-J. Li, *Tetrahedron Lett.*, 1997, **38**, 4731–4734.
2. M. A. Tekle-Smith, K. S. Williamson, I. F. Hughes and J. L. Leighton, *Org. Lett.*, 2017, **19**, 6024–6027.
3. J. Yuan, P. Jain and J. C. Antilla, *J. Org. Chem.*, 2022, **87**, 8256–8266.