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Supporting Information

Synthetic and Biological Studies on New Urea and Triazole Containing Cystobactamid Derivatives

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1. General information

Unless stated otherwise, all reactions conducted in organic solvents were performed in flame-dried glassware under an atmosphere of argon. Dry THF, DMF and CH_2Cl_2 were obtained by passing these previously degassed solvents through activated columns on a solvent purification system. Purchased reagents and chemicals were used as received, unless stated otherwise. Reactions were monitored by thin layer chromatography (TLC) on aluminum plates coated with silica gel, type 60 F254 by Merck and visualized by UV irradiation or development with a potassium permanganate or ninhydrine stain. Volatile solvents were removed under reduced pressure with a rotary evaporator. All column chromatographic purifications were performed using Machery-Nagel Silica 60 M (40 - 63 µm). Flash column chromatography was conducted with the flash purification system Sepacore® by Büchi using prepacked catridges (puriFlash® by Interchim or chromabond® by Machery-Nagel). Preparative high performance liquid chromatography using a Merck Hitachi LaChrom system (pump L- 7150, interface D-7000, diode array detector L-7450 (λ = 220-400 nm, preferred monitoring at λ = 230 nm)) with column (abbreviation referred to in the experimental part given in parentheses): Trentec Reprosil-Pur 120 C18 AQ 5 µm, 250 × 8 mm, with guard column, 40 × 8 mm (C18-SP) and a Thermo Scientific Dionex (Ultimate 300 HPLC system) 2 with a Kintex 5u Biphenyl 100A 250x10.0 mm column. ^{1}H-NMR and ^{13}C-NMR spectra were recorded with Bruker AVS or DRX spectrometers operating at 400, 500 or 600 MHz for ^{1}H (100, 125 or 150 MHz for ^{13}C) in DMSO-*d*$_6$. Chemical shifts are reported relative to the residual solvent signal (^{1}H-NMR: δ = 2.50 (DMSO-*d*$_6$); ^{13}C-NMR: δ = 39.5 (DMSO-*d*$_6$)). NMR data are reported as follows: chemical shift (δ) (multiplicity, coupling constants (J) where applicable, number of hydrogen atoms, assigned hydrogen). Splittings for ^{1}H-NMR signals are reported with the following symbols: bs = broad singlet, s = singlet, d = doublet, t = triplet, q = quartet, qi = quintet, h = heptet, dd = doublet of doublets, dt = doublet of triplets, dq = doublets of quartets, ddt = doublet of doublets of triplets, m = multiplet. For ^{13}C-NMR data the degree of substitution is reported as: p = primary C-atom, s = secondary C-atom, t = tertiary C-atom and q = quaternary C-atom. If necessary, COSY, HMBC and HSQC experiments were conducted for full characterization. High resolution mass spectra (HRMS) were obtained at 70 eV with a type VG Autospec spectrometer (Micromass), with a type LCT (ESI) (Micromass), with a type Q-TOF (Micromass) spectrometer in combination with a Waters Aquity Ultraperformance LC (UPLC) system, with a UPLC/Q-TOF-MS combination (Dionex Ultimate 3000/BrukerMaxis HD) or as gas chromatography with a HP6890. Optical rotations $[\alpha]^T_D$ were measured on Polarimeter 241 MC and 341 (both Perkin Elmer) at a wavelength of 589 nm (sodium

D line) and the concentration *c* is given in 10 mg mL^{-1} and the temperature in °C. Melting points were measured with OptiMelt (Stanford Research Systems). Melting points were measured using a SRS OptiMelt apparatus. 4-isocyanato-benzonitrile (**6**), 4-aminobenzoic acid (**7**), 1-isocyanto-4-nitrobenzene (**10**), 4-ethynylaniline (**29**), *tert*-butyl 4-amino-benzoate (**32**), 4-fluorobenzoyl chloride (**33**), 5-cyanopicolinic acid (**34**), 4-cyano-3-flurobenzoic acid (**35**), isonicotinic acid (**36**), 6-cyanonicotinic acid (**37**) and 3-(4-cyanophenyl)propanoic acid (**38**) are commercially available.

2. General procedures

General procedure I for amide formation (according to Scheme 1, main text)

DIPEA (4.0 eq) was added to a stirred solution of acid (2.0 eq), amine (1.0 eq) and COMU (2.0 eq) in DMF (0.15 M) at 0 °C. The mixture was stirred at rt for 15 h. The reaction was terminated by diluting with EtOAc, washed with sat. $NaHCO_3$ solution (4x), brine, dried over Na_2SO_4, filtered and concentrated under reduced pressure.

General procedure II for amide formation (according to Scheme 1, main text)

DIPEA (12.7 eq) was added dropwise to a stirred solution of acid (2.5 eq) and HATU (2.5 eq) in DMF (0.11 M) at rt and stirring was continued at rt for five minutes. The mixture was transferred dropwise to a stirred solution of amine (1.0 eq) in DMF (0.07 M) at rt and the mixture was stirred at rt for 15 h. The reaction was concentrated under reduced pressure.

General procedure III for amide formation (according to Scheme 1, main text)

Precooled chlouroform (0.33 M) was added to amine (1.0 eq) and acid (1.7 eq). A solution of EEDQ (1.6 eq) in precooled chloroform (0.75 M) was added dropwise to the acid-amine-mixture at 0 °C. After the mixture was warmed up to rt stirring was continued at rt for 16 h, it was dry loaded onto silica and purified by column chromatography (PE/EtOAc = 2:1 → 1:4).

General procedure IV for Boc deprotection (according to Scheme 1, main text)

Ester (1.0 eq) was added in small portions to 4 N HCl in 1,4-dioxane (100 eq) at 0 °C over five minutes, then the mixture was warmed up to rt over a period of 15 min. The mixture was slowly transferred to a mixture of EtOAc and a 80 % sat. aq. $NaHCO_3$ solution (400 mL, 1:1), the layers were separated, the aq. layer was extracted with EtOAc (3x 100 mL), the combined organic phases were washed with brine (150 mL), dried over Na_2SO_4, filtered and concentrated *in vacuo*.

General procedure V for *tert*-Bu ester hydrolysis (according to Scheme 1, main text)

Precooled TFA (0.02 M) was added slowly to ester (1.0 eq) at 0 °C. The mixture was warmed up to rt over a period 30 min, then it was stirred between 30 min and 6 h. The reaction was terminated by addition of Et_2O. The precipitate was filtered, washed with an excess of Et_2O and dried under high vacuum.

General procedure VI for de-*O*-allylation (according to Scheme 1, main text)

$Pd(PPh_3)_4$ (0.25 eq) was added in one portion to a stirred mixture of allyl ether (1.0 eq) and aniline (2.0 eq) in THF (0.04 M) and stirring was continued at rt for 2.5 h. The mixture was terminated by addition of HCl. The reaction was concentrated under reduced pressure.

3. Chemical Syntheses

4-(3-(4-Cyanophenyl)ureido)-benzoic acid (8): DIPEA (3.50 mL, 20.8 mmol, 3.0 eq) was added to a solution of 4-aminobenzoic acid (**7**, 952 mg, 6.94 mmol, 1.0 eq) and 4-cyanophenyl isocyanate (**6**, 1.00 g, 6.94 mmol, 1.0 eq) in THF (30 mL). The reaction was stirred at 55 °C for 16 h. The precipitate was filtered and washed with Et_2O (200 mL). Column chromatography (CH_2Cl_2/MeOH, 5-10% MeOH) yielded the title compound **8** (1.87 g, 6.66 mmol, 96%) as colourless solid.T_M = 290 °C (decomposition); ^{1}H-NMR (400 MHz, DMSO-d_6): δ = 12.7 (1H, br. s, CO_2*H*), 9.31 (1H, s, H-6/H-8), 9.23 (1H, s, H-6/H-8), 7.88 (2H, m, H-3, H-3‘), 7.74 (2H, m, H-10, H-10‘), 7.64 (2H, m, H-11, H-11‘), 7.57 (2H, m, H-4, H-4‘) ppm; ^{13}C-NMR (100 MHz, DMSO-d_6): δ = 167.0 (q, C-1), 151.9 (q, C-7), 143.9 (q, C-9), 143.4 (q, C-5), 133.3 (2C, t, C-11, C-11‘), 130.5 (2C, t, C-3, C-3‘), 124.2 (q, C-2), 119.2 (q, CN), 118.2 (2C, t, C-10, C-10‘), 117.6 (2C, t, C-4, C-4‘), 103.6 (q, C-12) ppm; HRMS (ESI): *m/z* calc. for $C_{15}H_{10}N_3O_3$ [M – H]$^-$: 280.0722; found 280.0723.

***tert*-Butyl-4-(2-(allyloxy)-3-isopropoxy-4-(3-(4-nitrophenyl)ureido)-benz-amido)benzoate (S1):** DIPEA (36 µL, 211 µmol, 2.0 eq) was added to a solution of amine **9** (45 mg, 106 µmol, 1.0 eq) and 4-nitrophenyl isocyanate (**10**, 52 mg, 317 µmol, 3.0 eq) in THF (2 mL). The reaction mixture was stirred at 55 °C for 6 h. Additional 4-nitrophenyl isocyanate (**9**, 52 mg, 317 µmol, 3.0 eq) was added and the reaction was stirred at 55 °C for another 14 h. The reaction mixture was concentrated *in vacuo* and the residue was purified by preparative HPLC (RP-18; run time 100 min; H_2O/MeCN = 70 : 30 → 0 : 100 in 80 min; t_r = 53 min) which provided compound **S1** as a colourless solid (44.7 mg, 76.4 mmol, 72%). T_M = 202 °C; ^{1}H-NMR (400 MHz, DMSO-d_6): δ = 10.4 (1H, s, H-8), 10.3 (1H, s, H-23), 8.55 (1H, s, H-21), 8.22 (2H, m, H-26, H-26‘), 8.09 (1H, d, J = 8.7 Hz, H-11), 7.88 (2H, m, H-5, H-5‘), 7.81 (2H, m, H-6, H-6‘), 7.73 (2H, m, H-25, H-25‘), 7.42 (1H, d, J = 8.7 Hz, H-12), 6.08 – 5.98 (1H, m, H-19), 5.40 (1H, dq, J = 1.6, 17.2 Hz, H-20$_{trans}$), 5.22 (1H, dq, J = 1.4, 10.4 Hz, H-20$_{cis}$), 4.66 – 4.59 (3H, m, H-16, H-18), 1.54 (9H, s, H-1), 1.33 (6H, d, J = 6.2 Hz, H-17) ppm; ^{13}C-NMR (100 MHz, DMSO-d_6):

δ = 164.6 (q, C-3), 164.3 (q, C-9), 151.6 (q, C-22), 149.3 (q, C-15), 146.0 (q, C-27), 143.0 (q, C-7), 141.3 (q, C-24), 138.9 (q, C-13), 136.8 (q, C-10), 133.6 (t, C-19), 130.1 (2C, t, C-5, C-5’), 126.0 (q, C-4), 125.3 (2C, t, C-26, C-26’), 124.4 (t, C-12), 124.1 (q, C-14), 118.8 (2C, t, C-6, C-6’), 117.9 (s, C-20), 117.6 (2C, t, C-25, C-25’), 114.0 (t, C-11), 80.3 (q, C-2), 76.2 (t, C-16), 74.2 (s, C-18), 27.9 (3C, p, C-1), 22.0 (2C, p, C-17) ppm; HRMS (ESI): *m/z* calc. for $C_{31}H_{34}N_4O_8Na$ $[M + Na]^+$: 613.2274; found 613.2278.

***tert*-Butyl-4-(2-(allyloxy)-4-(3-(4-aminophenyl)ureido)-3-isopropoxy-benzamido)benzoate (11):** Zinc (78.5 mg, 1.20 mmol, 2.0 eq) was added in one portion at rt to a stirred solution of nitro compound **S1** (352 mg, 0.60 mmol, 1.0 eq) in THF (1 mL), EtOH (1 mL) and glacial acetic acid (0.2 mL). After stirring vigorously at rt for 30 min, a 3rd and 4th equivalent of zinc were added and the mixture was stirred at rt for another 30 min. This step was repeated twice, after which time the reaction mixture was heated at 70 °C for a period of 30 min and the 9th and 10th equivalent of zinc were added and the mixture was stirred at 70 °C for additional 30 min. The reaction mixture was cooled to rt, diluted with Et_2O and terminated by addition of a sat. aq. $NaHCO_3$ solution. The precipitate was filtered and washed with an excess of Et_2O, the layers were separated, the aq. layer was extracted with Et_2O (3x), washed with brine, dried over Na_2SO_4, filtered and concentrated *in vacuo*. Column chromatography (PE/EtOAc= 3:1) yielded the title compound **11** (234 mg, 0.42 mmol, 70%) as a yellow solid. T_M = 134 °C; ^{1}H-NMR (400 MHz, DMSO-d_6): δ = 10.4 (1H, s, H-8), 9.05 (1H, s, H-21), 8.10 (1H, s, H-23), 8.08 (1H, s, H-11), 7.88 (2H, m, H-5, H-5‘), 7.80 (2H, m, H-6, H-6‘), 7.39 (1H, d, *J* = 8.8 Hz, H-12), 7.10 (2H, m, H-25, H-25‘), 6.53 (2H, m, H-26, H-26‘), 6.07 – 5.98 (1H, m, H-19), 5.40 (1H, dq, *J* = 1.6, 17.2 Hz, H-20_{trans}), 5.22 (1H, dq, *J* = 1.4, 9.3 Hz, H-20_{cis}), 4.83 (2H, br. S, NH_2), 4.62 – 4.57 (3H, m, H-16, H-18), 1.54 (9H, s, H-1), 1.29 (6H, d, *J* = 6.1 Hz, H-17) ppm; ^{13}C-NMR (100 MHz, DMSO-d_6): δ = 174.7 (q, C-3), 164.6 (q, C-9), 164.3 (q, C-22), 152.5 (q, C-13), 149.3 (q, C-15), 144.3 (q, C-27), 143.1 (q, C-7), 138.5 (q, C-10), 138.2 (q, C-14), 133.6 (t, C-19), 130.1 (2C, t, C-5, C-5’), 128.4 (q, C-24) 125.9 (q, C-4), 124.4 (t, C-12), 122.2 (2C, t, C-25, C-25’), 118.8 (2C, t, C-6, C-6’), 117.9 (s, C-20), 114.2 (2C, t, C-26, C-26’), 113.6 (t, C-11), 80.3 (q, C-2), 75.7 (t, C-16), 74.0 (s, C-18), 27.9 (3C, p, C-1), 21.9 (2C, p, C-17) ppm; HRMS (ESI): *m/z* calc. for $C_{31}H_{37}N_4O_6$ $[M + H]^+$: 561.2713; found 561.2705.

***tert*-Butyl-4-(2-(allyloxy)-3-isopropoxy-4-(4-nitrobenzamido)-benzamido)-benzoate (S2):** 4-Nitrobenzoyl chloride (3.48 g, 18.7 mmol, 1.6 eq) was added in small portions to a solution of amine **9** (5.00 g, 11.7 mmol, 1.0 eq) and pyridine (3.78 mL, 46.8 mmol, 4.0 eq) in CH_2Cl_2 (80 mL). The reaction was terminated after stirring 1.5 h at rt by addition of an aq. $NaHSO_4$ solution (1 M). The layers were separated and the aq. layer was extracted with CH_2Cl_2 (3x). The combined organic phases were dried over Na_2SO_4, filtered and concentrated *in vacuo* to yield crude nitro compound **S2** (6.66 g, 11.6 mmol, 99%) as a brown oil which was used without further purification in the next step.

***tert*-Butyl-4-(2-(allyloxy)-4-(4-aminobenzamido)-3-isopropoxybenzamido)-benzoate (12):** Zinc (757 mg, 11.6 mmol, 2.0 eq) was added in one portion at rt to a stirred solution of nitro compound **S2** (3.33 g, 5.79 mmol, 1.0 eq) in THF (9.7 mL), EtOH (9.7 mL) and glacial acetic acid (2.0 mL). After stirring vigorously at rt for 30 min, a 3rd and 4th equivalent of zinc were added and the mixture was stirred at rt for another 30 min. This step was repeated twice, after which time the reaction was heated to 70 °C for a period of 30 min and the 9th and 10th equivalent of zinc were added. The mixture was stirred at 70 °C for additional 30 min. The reaction mixture was cooled to rt, diluted with Et_2O and terminated by addition of a sat. aq. $NaHCO_3$ solution. The precipitate was filtered and washed with an excess of Et_2O, the layers were separated, the aq. layer was extracted with Et_2O (3x), washed with brine, dried over Na_2SO_4, filtered and concentrated *in vacuo*. Column chromatography (PE/EtOAc= 3:1) yielded the title compound **12** (2.59 g, 4.75 mmol, 82% over two steps) as a yellow solid. T_{M} = 90 °C; ^{1}H-NMR (400 MHz, $CDCl_3$): δ = 10.2 (1H, s, H-8), 8.65 (1H, s, H-21), 8.49 (1H, d, *J* = 8.9 Hz, H-11), 8.05 (1H, d, *J* = 8.9 Hz, H-12), 7.98 (2H, m, H-5, H-5‘), 7.74 (2H, m, H-24, H-24‘), 7.73 (2H, m, H-6, H-6‘), 6.73 (2H, m, H-25, H-25‘), 6.19 – 6.09 (1H, m, H-19), 5.49 (1H, dq, *J* = 1.4, 17.1 Hz, H-20$_{trans}$), 5.40 (1H, dq, *J* = 1.1, 10.4 Hz, H-20$_{cis}$), 4.73 (1H, hept, *J* = 6.2 Hz, H-16), 4.69 (2H, dt, *J* = 1.2, 5.9 Hz, H-18), 4.11 (2H, br. s, N*H*$_2$), 1.60 (9H, s, H-1), 1.38 (6H, d, *J* = 6.2 Hz, H-17) ppm; ^{13}C-NMR (100 MHz, $CDCl_3$): δ = 165.6 (q, C-3), 164.8 (q, C-22), 162.9 (q, C-9), 150.5 (q, C-26), 149.4 (q, C-15), 142.4 (q, C-7), 138.9 (q, C-14), 138.2 (q, C-4), 132.4 (t, C-19), 130.8 (2C, t, C-5, C-5’), 129.1 (2C, t, C-24, C-24’), 127.6 (q, C-13), 127.3 (t, C-12), 123.8 (q, C-23), 121.1 (q, C-10), 120.1 (s, C-20), 119.1 (2C, t, C-6, C-6’), 115.7 (t, C-11), 114.5 (2C, t, C-25, C-25’), 80.9 (q, C-2), 76.7 (t, C-16), 75.0 (s, C-18), 28.4 (3C, p, C-1), 23.0 (2C, p, C-17) ppm; HRMS (ESI): *m/z* calc. for $C_{31}H_{36}N_3O_6$ $[M + H]^+$: 546.2604; found 546.2600.

***tert*-Butyl-4-(2-(allyloxy)-4-(4-((2*S*,3*R*)-4-amino-2-((*tert*-butoxycarbonyl)-amino)-3-methoxy-4-oxobutanamido)benzamido)-3-isopropoxybenz-amido)benzoate (S3):** Following to the general procedure III using amine **12** (1.50 g, 2.75 mmol, 1.0 eq) and acid **13** (1.23 g, 4.68 mmol, 1.7 eq) compound **S3** (1.37 g, 1.73 mmol, 63%) was obtained as a yellow amorphous solid. $[\alpha]_D^{22}$ = -0.9° (*c* 1.3, MeOH); ^{1}H-NMR (400 MHz, DMSO-d_6): δ = 10.5 (1H, s, N*H*), 10.4 (1H, s, N*H*), 9.51 (1H, s, N*H*), 7.98 (2H, m, Ar*H*), 7.90 (2H, m, Ar*H*), 7.84 – 7.80 (5H, m, Ar*H*), 7.45 – 7.40 (3H, m, Ar*H*, N*H*$_2$), 6.80 (1H, d, *J* = 8.4 Hz, N*H*Boc), 6.07 – 5.97 (1H, m, C*H*$_{\text{allyl}}$), 5.37 (1H, dd, *J* = 1.6, 17.1 Hz, C*H*$_{\text{allyl-}trans}$), 5.20 (1H, dd, *J* = 1.4, 9.8 Hz, C*H*$_{\text{allyl-}cis}$), 4.61 (2H, d, *J* = 5.4 Hz, OC*H*$_2$allyl), 4.50 (1H, hept, *J* = 6.1 Hz, C*H*Me$_2$), 4.41 (1H, t, *J* = 7.9 Hz, C*H*NH), 3.86 (1H, d, *J* = 7.4 Hz, C*H*OMe), 3.26 (3H, s, OCH_3), 1.55 (9H, s, C(CH_3)$_3$), 1.38 (9H, s, C(CH_3)$_3$), 1.26 (6H, d, *J* = 6.2 Hz, CH(CH_3)$_2$) ppm; ^{13}C-NMR (100 MHz, DMSO-d$_6$): δ = 170.6 (q, CONH$_2$), 168.9 (q, *C*ONH), 164.6 (q, *C*ONH), 164.5 (q, *C*ONH), 164.3 (q, CO$_2$*t*Bu), 154.8 (q, CO$_2$*t*Bu), 149.5 (q, C-Ar), 143.0 (q, C-Ar), 142.5 (q, C-Ar), 142.2 (q, C-Ar), 135.7 (q, C-Ar), 133.6 (t, *C*Hallyl), 130.1 (2C, t, C-Ar), 128.4 (2C, t, C-Ar), 127.0 (q, C-Ar), 126.0 (q, C-Ar), 123.6 (t, C-Ar), 118.9 (t, C-Ar), 118.8 (4C, t, C-Ar), 117.8 (s, *C*H$_2$allyl), 80.3 (t, *C*HOMe), 80.1 (q, *C*(CH$_3$)$_3$), 78.7 (q, *C*(CH$_3$)$_3$), 76.3 (t, *C*H(CH$_3$)$_2$), 74.3 (s, OCH_2allyl), 57.6 (p, O*C*H$_3$), 56.6 (t, *C*HNH), 28.1 (3C, p, C(*C*H$_3$)$_3$), 27.8 (3C, p, C(*C*H$_3$)$_3$), 22.3 (2C, p, CH(*C*H$_3$)$_2$) ppm; HRMS (ESI): *m/z* calc. for $C_{41}H_{51}N_5O_{11}Na$ [M + Na]$^+$: 812.3483; found 812.3480.

***tert*-Butyl-4-(2-(allyloxy)-4-(4-((2*S*,3*R*)-2,4-diamino-3-methoxy-4-oxobutan-amido)-benzamido)-3-isopropoxybenzamido)benzoate (14):** Following the general procedure IV with **S3** (200 mg, 0.25 mmol, 1.0 eq) preparative HPLC (RP-18; run time 100 min; H_2O/MeCN = 80 : 20 → 0 : 100 in 80 min; t$_r$ = 32 min) provided amine **14** as a yellow amorphous solid (83 mg, 0.12 mmol, 48%). $[\alpha]_D^{23}$ = +3.8° (*c* 1.1, MeOH); ^{1}H-NMR (400 MHz, DMSO-d$_6$): δ = 10.5 (1H, s, NH), 9.53 (1H, s, NH), 7.98 (2H, m, ArH), 7.89 (2H, m, ArH), 7.84 – 7.80 (5H, m, ArH), 7.47 (2H, d, *J* = 70 Hz, NH$_2$), 7.41 (1H, d, *J* = 8.5 Hz, ArH), 6.07 – 5.97 (1H, m, CH_{allyl}), 5.38 (1H, dd, *J* = 1.7, 17.1 Hz, C$H_{\text{allyl-}trans}$), 5.20 (1H, dd, *J* = 1.5, 9.8 Hz, C$H_{\text{allyl-}cis}$), 4.61 (2H, d, *J* = 5.5 Hz, OCH_2allyl), 4.50 (1H, hept, *J* = 6.1 Hz, CHMe$_2$), 3.86 (1H, d, *J* = 6.6 Hz, CHOMe), 3.79 (1H, d, *J* = 6.6 Hz, CHNH$_2$), 3.31 (3H, s, OCH_3), 1.55 (9H, s, C(CH_3)$_3$), 1.26 (6H, d, *J* = 6.1 Hz, CH(CH_3)$_2$) ppm; ^{13}C-NMR (100 MHz, DMSO-d$_6$): δ = 171.1 (q, CONH$_2$), 164.6 (q, *C*ONH), 164.5 (q, *C*ONH), 164.3 (q, *C*ONH), 163.0 (q, CO$_2$*t*Bu), 149.5 (q, C-Ar), 143.0 (q, C-Ar), 142.5 (q, C-Ar), 142.0 (q, C-Ar), 135.6 (q, C-Ar), 133.6 (t, *C*Hallyl), 130.1 (2C, t, C-Ar), 128.5 (q, C-Ar), 128.4 (2C, t, C-Ar), 127.1 (q, C-Ar), 126.0 (q, C-Ar), 123.6 (t, C-Ar), 119.0 (t, C-Ar), 118.8 (2C, t, C-Ar), 118.8 (2C, t, C-Ar), 117.8 (s, *C*H$_2$allyl), 82.1 (t, *C*HOMe), 80.3 (q, *C*(CH$_3$)$_3$), 76.2 (t, *C*H(CH$_3$)$_2$), 74.3 (s, OCH_2allyl), 58.0 (p, O*C*H$_3$), 56.6 (t, *C*HNH$_2$), 27.9 (3C, p, C(*C*H$_3$)$_3$), 22.3 (2C, p, CH(*C*H$_3$)$_2$) ppm; HRMS (ESI): *m/z* calc. for $C_{36}H_{43}N_5O_9Na$ [M + Na]$^+$: 712.2958; found 712.2960.

***tert*-Butyl-4-(2-(allyloxy)-4-(4-((2*S*,3*R*)-4-amino-2-(4-(3-(4-cyanophenyl)-ureido)-benz-amido)-3-methoxy-4-oxobutanamido)benzamido)-3-iso-propoxybenzamido)benzoate (15):** Following the general procedure II acid **8** (41.8 mg, 149 µmol, 2.5 eq) and amine **14** (41.0 mg, 59.4 µmol, 1.0 eq). provided compound **15** as a colourless solid after preparative HPLC (RP-18; run time 100 min; H_2O/MeCN = 60 : 40 → 0 : 100 in 80 min; t$_r$ = 67 min) (42.7 mg, 44.8 µmol, 75%).T$_M$ = 226 °C (decomposition); $[\alpha]_D^{22}$ = +34.0° (*c* 4.2, MeOH); ^{1}H-NMR (400 MHz, DMSO-d$_6$): δ = 10.5 (1H, s, NH), 10.5 (1H, s, NH), 9.52 (1H, s, NH), 9.51 (1H, s, NH_{Urea}), 9.38 (1H, s, NH_{Urea}), 8.37 (1H, d, *J* = 8.1 Hz, NHCH), 7.98 (2H, m, ArH), 7.89 (2H, m, ArH), 7.84 – 7.82 (7H, m, ArH), 7.74 (2H, m, ArH), 7.66

(2H, m, Ar*H*), 7.59 (2H, m, Ar*H*), 7.51 (2H, d, *J* = 27.1 Hz, CON*H*$_2$), 7.41 (1H, d, *J* = 8.5 Hz, Ar*H*), 6.07 – 5.97 (1H, m, C*H*$_{\text{allyl}}$), 5.38 (1H, dq, *J* = 1.6, 17.2 Hz, C*H*$_{\text{allyl-}\textit{trans}}$), 5.20 (1H, dq, *J* = 1.3, 10.5 Hz, C*H*$_{\text{allyl-}\textit{cis}}$), 4.90 (1H, t, *J* = 8.1 Hz, C*H*NH), 4.61 (2H, d, *J* = 5.4 Hz, OC*H*$_2$allyl), 4.50 (1H, hept, *J* = 6.1 Hz, C*H*Me$_2$), 4.09 (1H, d, *J* = 8.1 Hz, C*H*OMe), 3.01 (3H, s, OC*H*$_3$), 1.54 (9H, s, C(C*H*$_3$)$_3$), 1.26 (6H, d, *J* = 6.1 Hz, CH(C*H*$_3$)$_2$) ppm; ^{13}C-NMR (100 MHz, DMSO-d$_6$): δ = 170.9 (q, CONH$_2$), 168.8 (q, *C*ONH), 165.5 (q, *C*ONH), 164.6 (q, *C*ONH), 164.5 (q, CO$_2$*t*Bu), 164.3 (q, *C*ONH), 152.0 (q, NH*C*ONH), 149.5 (q, C-Ar), 144.0 (q, C-Ar), 143.0 (q, C-Ar), 142.5 (q, C-Ar), 142.5 (q, C-Ar), 142.2 (q, C-Ar), 135.7 (q, C-Ar), 133.6 (t, *C*Hallyl), 133.3 (2C, t, C-Ar), 130.1 (2C, t, C-Ar), 128.5 (2C, t, C-Ar), 128.5 (t, C-Ar), 128.4 (2C, t, C-Ar), 127.1 (q, C-Ar), 126.0 (q, C-Ar), 123.6 (t, C-Ar), 119.3 (q, CN), 118.9 (2C, t, C-Ar), 118.8 (2C, t, C-Ar), 118.2 (2C, t, C-Ar), 117.8 (s, *C*H$_2$allyl), 117.5 (2C, t, C-Ar), 103.5 (q, C-Ar), 80.3 (q, *C*(CH$_3$)$_3$), 80.0 (t, *C*HOMe), 76.3 (t, *C*H(CH$_3$)$_2$), 74.3 (s, OC*H*$_2$allyl), 57.7 (p, O*C*H$_3$), 55.7 (t, *C*HNH), 27.9 (3C, p, C(*C*H$_3$)$_3$), 22.3 (2C, p, CH(*C*H$_3$)$_2$) ppm; HRMS (ESI): *m/z* calc. for $C_{51}H_{52}N_8O_{11}Na$ [M + Na]$^+$: 975.3653; found 975.3633.

***tert*-Butyl-4-(4-(4-((2*S*,3*R*)-4-amino-2-(4-(3-(4-cyanophenyl)ureido)-benz-amido)-3-methoxy-4-oxobutanamido)benzamido)-2-hydroxy-3-isopropoxy-benzamido)benzoate (S4):** Following to the general procedure VI compound **15** (42.0 mg, 44.1 µmol, 1.0 eq) was used. Preparative HPLC (RP-18; run time 80 min; H$_2$O/MeCN = 70 : 30 → 0 : 100 in 70 min; t$_r$ = 56 min) provided Phenol **S4** as a colourless amorphous solid (29.8 mg, 32.6 µmol, 74%). $[\alpha]_D^{21}$ = +29.3° (*c* 1.2, MeOH); ^{1}H-NMR (400 MHz, DMSO-d$_6$): δ = 12.4 (1H, br. s, O*H*), 10.8 (1H, br. s, N*H*), 10.6 (1H, s, N*H*), 9.40 (1H, s, N*H*), 9.38 (1H, s, N*H*$_{\text{Urea}}$), 9.28 (1H, s, N*H*$_{\text{Urea}}$), 8.38 (1H, d, *J* = 8.1 Hz, N*H*CH), 7.96 (2H, m, Ar*H*), 7.92 (2H, m, Ar*H*), 7.86 – 7.82 (7H, m, Ar*H*), 7.74 (2H, m, Ar*H*), 7.70 (1H, d, *J* = 8.9 Hz, Ar*H*), 7.66 (2H, m, Ar*H*), 7.59 (2H, m, Ar*H*), 7.51 (2H, d, *J* = 26.5 Hz, CON*H*$_2$), 4.91 (1H, t, *J* = 8.1 Hz, C*H*NH), 4.56 (1H, hept, *J* = 6.2 Hz, C*H*Me$_2$), 4.09 (1H, d, *J* = 8.1 Hz, C*H*OMe), 3.31 (3H, s, OC*H*$_3$), 1.55 (9H, s, C(C*H*$_3$)$_3$), 1.27 (6H, d, *J* = 6.2 Hz, CH(C*H*$_3$)$_2$) ppm; ^{13}C-NMR (100 MHz, DMSO-d$_6$): δ = 170.9 (q, CONH$_2$), 168.8 (q, *C*ONH), 168.4 (q, *C*ONH), 165.5 (q, *C*ONH), 164.6 (q, CO$_2$*t*Bu), 164.2 (q, *C*ONH), 154.5 (q, C-Ar), 152.0 (q, NH*C*ONH), 144.0 (q, C-Ar), 142.4 (q, C-Ar), 142.3 (q, C-Ar), 142.1 (q, C-Ar), 137.0 (q, C-Ar), 136.3 (q, C-Ar), 133.3 (2C, t, C-Ar), 129.9 (2C, t, C-Ar), 128.5 (2C, t, C-Ar), 128.3 (2C, t, C-Ar), 127.1 (q, C-Ar), 126.7 (q, C-Ar), 122.9 (t, C-Ar), 120.6 (2C, t, C-Ar), 119.3 (q, CN), 119.0 (2C, t, C-Ar), 118.2 (2C, t, C-Ar), 117.5 (2C, t, C-Ar), 112.5 (t, C-Ar), 111.8 (q, C-Ar), 103.5 (q, C-Ar), 80.5 (q, *C*(CH$_3$)$_3$), 80.0 (t, *C*HOMe), 74.7 (t, *C*H(CH$_3$)$_2$), 57.7 (p, O*C*H$_3$), 55.8 (t, *C*HNH), 27.9 (3C, p, C(*C*H$_3$)$_3$), 22.3 (2C, p, CH(*C*H$_3$)$_2$) ppm; HRMS (ESI): *m/z* calc. for $C_{48}H_{48}N_8O_{11}Na$ [M + Na]$^+$: 935.3340; found 935.3342.

4-(4-(4-((2*S*,3*R*)-4-Amino-2-(4-(3-(4-cyanophenyl)ureido)benzamido)-3-methoxy-4-oxobutanamido)benzamido)-2-hydroxy-3-isopropoxybenz-amido)benzoic acid (16): Following to the general procedure V phenol **S4** (29.8 mg, 32.6 µmol, 1.0 eq) was stirred for 5.5 h. Acid **16** was obtained as a colourless amorphous solid (22.5 mg, 26.3 µmol, 81%). $[\alpha]_D^{22}$ = +23.1° (*c* 1.1, MeOH); ^{1}H-NMR (400 MHz, DMSO-d$_6$): δ = 12.3 (1H, br. s, O*H*), 10.6 (1H, br. s, N*H*), 10.6 (1H, s, N*H*), 9.40 (1H, s, N*H*), 9.38 (1H, s, N*H*$_{\text{Urea}}$), 9.26 (1H, s, N*H*$_{\text{Urea}}$), 8.37 (1H, d, *J* = 8.1 Hz, N*H*CH), 7.98 – 7.95 (4H, m, Ar*H*), 7.88 – 7.82 (7H, m, Ar*H*), 7.76 – 7.70 (3H, m, Ar*H*), 7.66 (2H, m, Ar*H*), 7.58 (2H, m, Ar*H*), 7.51 (2H, d, *J* = 26.7 Hz, CON*H*$_2$), 4.90 (1H, t, *J* = 8.1 Hz, C*H*NH), 4.55 (1H, hept, *J* = 6.1 Hz, C*H*Me$_2$), 4.09 (1H, d, *J* = 8.1 Hz, C*H*OMe), 3.31 (3H, s, OC*H*$_3$), 1.27 (6H, d, *J* = 6.1 Hz, CH(C*H*$_3$)$_2$) ppm; ^{13}C-NMR (100 MHz, DMSO-d$_6$): δ = 170.9 (q, CONH$_2$), 168.8 (q, *C*ONH), 168.5 (q, *C*ONH), 166.9 (q, *C*O$_2$H), 165.5 (q, *C*ONH), 164.2 (q, *C*ONH), 154.1 (q, C-Ar), 151.9 (q, NH*C*ONH), 144.0 (q, C-Ar), 142.4 (q, C-Ar), 142.3 (q, C-Ar), 142.0 (q, C-Ar), 137.0 (q, C-Ar), 136.3 (q, C-Ar), 133.3 (2C, t, C-Ar), 130.2 (2C, t, C-Ar), 128.5 (2C, t, C-Ar), 128.5 (q, C-Ar), 128.3 (2C, t, C-Ar), 127.1 (q, C-Ar), 126.3 (q, C-Ar), 122.8 (t, C-Ar), 120.7 (2C, t, C-Ar), 119.3 (q, CN), 119.0 (2C, t, C-Ar), 118.2 (2C, t, C-Ar), 117.5 (2C, t, C-Ar), 112.4 (t, C-Ar), 112.2 (q, C-Ar), 103.5 (q, C-Ar), 80.0 (t, *C*HOMe), 74.9 (t, *C*H(CH$_3$)$_2$), 57.7 (p, O*C*H$_3$), 55.7 (t, *C*HNH), 22.3 (2C, p, CH(*C*H$_3$)$_2$) ppm; HRMS (ESI): *m/z* calc. for $C_{44}H_{41}N_8O_{11}$ [M + H]$^+$: 857.2895; found 857.2896.

***tert*-Butyl-4-(2-(allyloxy)-4-(3-(4-((2*S*,3*R*)-4-amino-2-((*tert*-butoxycarbonyl)-amino)-3-methoxy-4-oxobutanamido)phenyl)ureido)-3-isopropoxybenz-amido)benzoate (17):** Following the general procedure III using amine **11** (234 mg, 417 µmol, 1.0 eq) and acid **13** (186 mg, 710 µmol, 1.7 eq) compound **17** (229 mg, 284 mmol, 68%) was obtained as a yellow solid. T_M = 168 °C; $[\alpha]_D^{21}$ = -7.0° (*c* 0.9, MeOH); ^{1}H-NMR (400 MHz, DMSO-d$_6$): δ = 10.4 (1H, s, N*H*), 9.94 (1H, s, N*H*), 9.52 (1H, s, N*H*$_{\text{Urea}}$), 8.30 (1H, s, N*H*$_{\text{Urea}}$), 8.11 (1H, d, *J* = 8.8 Hz, Ar*H*), 7.88 (2H, m, Ar*H*), 7.81 (2H, m, Ar*H*), 7.57 (2H, m, Ar*H*), 7.42 – 7.38 (5H, m, Ar*H*, N*H*$_2$), 6.70 (1H, d, *J* = 8.3 Hz, N*H*Boc), 6.05 – 5.98 (1H, m, C*H*$_{\text{allyl}}$), 5.40 (1H, dd, *J* = 1.6, 17.2 Hz, C*H*$_{\text{allyl-}\textit{trans}}$), 5.22 (1H, dd, *J* = 1.4, 9.3 Hz, C*H*$_{\text{allyl-}\textit{cis}}$), 4.64 – 4.58 (3H, m, OC*H*$_2$allyl, C*H*Me$_2$), 4.35 (1H, t, *J* = 8.5 Hz, C*H*NH), 3.83 (1H, d, *J* = 7.4 Hz, C*H*OMe), 3.25 (3H, s, OC*H*$_3$), 1.54 (9H, s, C(C*H*$_3$)$_3$), 1.38 (9H, s, C(C*H*$_3$)$_3$), 1.33 (6H, d, *J* = 6.1 Hz, CH(C*H*$_3$)$_2$) ppm; ^{13}C-NMR (100 MHz, DMSO-d$_6$): δ = 170.8 (q, CONH$_2$), 167.8 (q, *C*ONH), 164.6 (2C, q, *C*ONH), 164.3 (q, CO$_2$*t*Bu), 152.0 (q, NH*C*ONH), 149.3 (q, C-Ar), 143.1 (q, C-Ar), 138.4 (q, C-Ar), 137.8 (q, C-Ar), 135.0 (q, C-Ar), 133.6 (t, *C*Hallyl), 130.1 (2C, t, C-Ar), 125.9 (q, C-Ar), 124.5 (t, C-Ar), 124.4 (q, C-Ar), 123.0 (q, C-Ar), 120.1 (2C, t, C-Ar), 118.8 (2C, t, C-Ar), 118.6 (2C, t, C-Ar), 117.9 (s, *C*H$_2$allyl), 113.7 (t, C-Ar), 80.3 (t, *C*HOMe), 78.6 (2C, q, *C*(CH$_3$)$_3$), 75.9 (t, *C*H(CH$_3$)$_2$), 74.1 (s, OC*H*$_2$allyl), 57.6 (p, O*C*H$_3$), 56.5 (t, *C*HNH), 28.1 (3C, p, C(*C*H$_3$)$_3$), 27.9 (3C, p, C(*C*H$_3$)$_3$), 21.9 (2C, p, CH(*C*H$_3$)$_2$) ppm; **HRMS** (ESI): *m/z* calc. for $C_{41}H_{53}N_6O_{11}$ [M + H]$^+$: 805.3772; found 805.3772.

***tert*-Butyl-4-(2-(allyloxy)-4-(3-(4-((2*S*,3*R*)-2,4-diamino-3-methoxy-4-oxobutanamido)-phenyl)ureido)-3-isopropoxybenzamido)benzoate (18):** Following to the general procedure IV compound **17** (178 mg, 0.22 mmol, 1.0 eq) was used. Preparative HPLC (RP-18; run time 100 min; H_2O/MeCN = 70 : 30 → 0 : 100 in 80 min; t_r = 26 min) provided amine **18** as a yellow amorphous solid (76.2 mg, 0.11 mmol, 49%). $[\alpha]_D^{21}$ = +4.1° (*c* 1.1, MeOH); ^{1}H-NMR (400 MHz, DMSO-d_6): δ = 10.4 (1H, s, N*H*), 9.87 (1H, br. s, N*H*), 9.59 (1H, s, N*H*$_{\text{Urea}}$), 8.33 (1H, s, N*H*$_{\text{Urea}}$), 8.11 (1H, d, *J* = 8.8 Hz, Ar*H*), 7.88 (2H, m, Ar*H*), 7.81 (2H, m, Ar*H*), 7.58 (2H, m, Ar*H*), 7.43 – 7.41 (3H, m, Ar*H*), 7.36 (2H, d, *J* = 80.3 Hz, CON*H*$_2$), 6.07 – 5.98 (1H, m, C*H*$_{\text{allyl}}$), 5.40 (1H, dd, *J* = 1.6, 17.2 Hz, C*H*$_{\text{allyl-}trans}$), 5.22 (1H, dd, *J* = 1.4, 9.3 Hz, C*H*$_{\text{allyl-}cis}$), 4.65 – 4.59 (3H, m, OC*H*$_2$allyl, C*H*Me$_2$), 3.76 (1H, d, *J* = 6.8 Hz, C*H*OMe), 3.59 (1H, d, *J* = 6.8 Hz, C*H*NH$_2$), 3.37 (2H, br. s, N*H*$_2$), 3.28 (3H, s, OC*H*$_3$), 1.54 (9H, s, C(C*H*$_3$)$_3$), 1.33 (6H, d, *J* = 6.1 Hz, CH(C*H*$_3$)$_2$) ppm; ^{13}C-NMR (100 MHz, DMSO-d_6): δ = 171.7 (q, CONH$_2$), 170.7 (q, *C*ONH), 164.6 (q, CO$_2$*t*Bu), 164.3 (q, *C*ONH), 152.1 (q, NH*C*ONH), 149.3 (q, C-Ar), 143.1 (q, C-Ar), 138.4 (q, C-Ar), 137.8 (q, C-Ar), 134.9 (q, C-Ar), 133.7 (q, C-Ar), 133.6 (t, *C*Hallyl), 130.1 (2C, t, C-Ar), 125.9 (q, C-Ar), 124.5 (t, C-Ar), 122.9 (q, C-Ar), 119.9 (2C, t, C-Ar), 118.8 (2C, t, C-Ar), 118.7 (2C, t, C-Ar), 117.9 (s, *C*H$_2$allyl), 113.7 (t, C-Ar), 83.2 (t, *C*HOMe), 80.3 (q, *C*(CH$_3$)$_3$), 75.9 (t, *C*H(CH$_3$)$_2$), 74.1 (s, OC*H*$_2$allyl), 57.8 (p, O*C*H$_3$), 57.1 (t, *C*HNH$_2$), 27.9 (3C, p, C(*C*H$_3$)$_3$), 21.9 (2C, p, CH(*C*H$_3$)$_2$) ppm; HRMS (ESI): *m/z* calc. for $C_{36}H_{45}N_6O_9$ $[M + H]^+$: 705.3248; found 705.3242.

4-(2-(Allyloxy)-4-(3-(4-((2*S*,3*R*)-2,4-diamino-3-methoxy-4-oxobutanamido)-phenyl)-ureido)-3-isopropoxybenzamido)benzoic acid (S5): Ester **17** (80 mg, 99.4 µmol, 1.0 eq) was added in small portions to 4 N HCl in 1,4-dioxane (2.48 mL, 9.94 mmol, 100 eq) and stirred for 30 min at rt. The mixture was slowly transferred to a mixture of EtOAc and a 80 % sat. aq. $NaHCO_3$ solution (400 mL, 1:1). The layers were separated, the aq. layer was extracted with EtOAc (3x 100 mL), the combined organic phases were washed with brine (150 mL), dried over Na_2SO_4, filtered and concentrated *in vacuo*. Preparative HPLC (RP-18; run time 100 min; H_2O/MeCN = 95 : 05 → 0 : 100 in 90 min; t_r = 30 min) provided acid **S5** as a yellow amorphous solid (57.3 mg, 88.4 µmol, 89%). $[\alpha]_D^{22}$ = +7.0° (*c* 0.9, MeOH); ^{1}H-NMR (400 MHz, DMSO-d_6): δ = 10.4 (1H, s, N*H*), 9.87 (1H, br. s, N*H*), 9.56 (1H, s, N*H*$_{\text{Urea}}$), 8.31 (1H, s, N*H*$_{\text{Urea}}$), 8.11 (1H, d, *J* = 8.8 Hz, Ar*H*), 7.92 (2H, m, Ar*H*), 7.81 (2H, m, Ar*H*), 7.58 (2H, m, Ar*H*), 7.42 – 7.40 (3H, m, Ar*H*), 7.37 (2H, d, *J* = 80.7 Hz, CON*H*$_2$), 6.08 – 5.99 (1H, m, C*H*$_{\text{allyl}}$), 5.40 (1H, dd, *J* = 1.7, 17.2 Hz, C*H*$_{\text{allyl-}trans}$), 5.23 (1H, dd, *J* = 1.4, 10.4 Hz, C*H*$_{\text{allyl-}cis}$), 4.64 – 4.59 (3H, m, OC*H*$_2$allyl, C*H*Me$_2$), 3.76 (1H, d, *J* = 6.8 Hz, C*H*OMe), 3.60 (1H, d, *J* = 6.8 Hz, C*H*NH$_2$), 3.27 (3H, s, OC*H*$_3$), 1.33 (6H, d, *J* = 6.1 Hz, CH(C*H*$_3$)$_2$) ppm; ^{13}C-NMR (100 MHz, DMSO-d_6): δ = 171.7 (q, *C*ONH$_2$), 170.7 (q, *C*ONH), 167.1 (q, *C*O$_2$H), 164.3 (q, *C*ONH), 152.1 (q, NH*C*ONH), 149.3 (q, C-Ar), 142.9 (q, C-Ar), 138.5 (q, C-Ar), 137.8 (q, C-Ar), 134.9 (q, C-Ar), 133.7 (q, C-Ar), 133.6 (t, *C*Hallyl), 130.4 (2C, t, C-Ar), 125.8 (q, C-Ar), 124.5 (t, C-Ar), 123.1 (q, C-Ar), 119.9 (2C, t, C-Ar), 118.9 (2C, t, C-Ar), 118.7 (2C, t, C-Ar), 117.9 (s, *C*H$_2$allyl), 113.7 (t, C-Ar), 83.2 (t, *C*HOMe), 76.0 (t, *C*H(CH$_3$)$_2$), 74.1 (s, OC*H*$_2$allyl), 57.8 (p, O*C*H$_3$), 57.1 (t, *C*HNH$_2$), 22.0 (2C, p, CH(*C*H$_3$)$_2$) ppm; HRMS (ESI): *m/z* calc. for $C_{32}H_{37}N_6O_9$ $[M + H]^+$: 649.2622; found 649.2624.

4-(4-(3-(4-((2*S*,3*R*)-2,4-diamino-3-methoxy-4-oxobutanamido)phenyl)-ureido)-2-hydroxy-3-isopropoxybenzamido)benzoic acid (25): Following to the general procedure VI compound **S5** (10.0 mg, 15.4 µmol, 1.0 eq) provided phenol **25** as a colourless amorphous solid (6.5 mg, 10.7 µmol, 69%). Preparative HPLC (RP-18; run time 80 min; H$_2$O/MeCN = 95 : 05 → 0 : 100 in 60 min; t_r = 21 min). $[\alpha]_D^{23}$ = +24.1° (*c* 0.7, MeOH); ^{1}H-NMR (500 MHz, DMSO-d$_6$): δ = 9.83 (1H, br. s, CO$_2$*H*), 9.55 (1H, s, N*H*), 8.34 (2H, br. s, N*H*$_{\text{Urea}}$), 8.09 (1H, s, N*H*), 7.87 (2H, m, Ar*H*), 7.76 (2H, m, Ar*H*), 7.57 – 7.54 (3H, m, Ar*H*), 7.46 – 7.43 (2H, m, Ar*H*, CON*H*$_2$), 7.40 (2H, m, Ar*H*), 7.24 (1H, s, CON*H*$_2$), 4.79 (1H, hept, *J* = 6.5 Hz, C*H*Me$_2$), 3.74 (1H, d, *J* = 7.0 Hz, C*H*OMe), 3.57 (1H, d, *J* = 7.0 Hz, C*H*NH$_2$), 3.27 (3H, s, OC*H*$_3$), 1.25 (6H, d, *J* = 6.5 Hz, CH(C*H*$_3$)$_2$) ppm; ^{13}C-NMR (125 MHz, DMSO-d$_6$): δ = 171.8 (q, *C*ONH$_2$), 170.8 (q, *C*ONH), 168.0 (q, *C*O$_2$H), 164.8 (q, *C*ONH), 152.2 (q, NH*C*ONH), 143.5 (q, C-Ar), 137.4 (q, C-Ar), 135.3 (q, C-Ar), 134.5 (q, C-Ar), 133.9 (q, C-Ar), 133.9 (q, C-Ar), 130.4 (q, C-Ar), 130.2 (2C, t, C-Ar), 123.4 (t, C-Ar), 119.8 (2C, t, C-Ar), 119.2 (2C, t, C-Ar), 118.5 (2C, t, C-Ar), 112.0 (q, C-Ar), 104.0 (t, C-Ar), 83.3 (t, *C*HOMe), 72.0 (t, *C*H(CH$_3$)$_2$), 57.7 (p, O*C*H$_3$), 57.1 (t, *C*HNH$_2$), 22.1 (2C, p, CH(*C*H$_3$)$_2$) ppm; HRMS (ESI): *m/z* calc. for $C_{29}H_{32}N_6O_9Na$ $[M + Na]^+$: 631.2128; found 631.2127.

***tert*-Butyl-4-(2-(allyloxy)-4-(3-(4-((2*S*,3*R*)-4-amino-3-methoxy-2-(4-(4-nitro-benzamido)-benzamido)-4-oxobutanamido)phenyl)ureido)-3-isopropoxy-benzamido)benzoate (21):** Following the general procedure II acid **19** (31.0 mg, 108 µmol, 2.5 eq) and amine **18** (31.0 mg, 44.0 µmol, 1.0 eq) provided phenol **S7** as a grey amorphous solid (15.8 mg, 17.3 µmol, 61%). Preparative HPLC (RP-18; run time 100 min; H$_2$O/MeCN = 70 : 30 → 0 : 100 in 80 min; t_r = 57 min) provided compound **21** as a colourless solid (25.2 mg, 25.9 µmol, 59%). T_M = 230 °C (decomposition); $[\alpha]_D^{21}$ = +46.1° (*c* 0.9, MeOH); ^{1}H-NMR (400 MHz, DMSO-d$_6$): δ = 10.8 (1H, s, N*H*), 10.4 (1H, s, N*H*), 10.2 (1H, s, N*H*), 9.67 (1H, s, N*H*$_{\text{Urea}}$), 8.43 – 8.35 (4H, m, Ar*H*, N*H*$_{\text{Urea}}$, N*H*CH), 8.22 (2H, m, Ar*H*), 8.11 (1H, d, *J* = 8.7 Hz, Ar*H*), 7.91 – 7.87 (6H, m, Ar*H*), 7.81 (2H, m, Ar*H*), 7.61 – 7.55 (3H, m, Ar*H*, CON*H*$_2$), 7.44 – 7.40 (4H, m, Ar*H*, CON*H*$_2$), 6.06 – 5.98 (1H, m, C*H*$_{\text{allyl}}$), 5.40 (1H, dd, *J* = 1.4, 17.3 Hz, C*H*$_{\text{allyl-}trans}$), 5.22 (1H, dd, *J* = 1.1, 9.2 Hz, C*H*$_{\text{allyl-}cis}$), 4.89 (1H, t, *J* = 8.1 Hz, C*H*NH), 4.63 – 4.59 (3H, m, OC*H*$_2$allyl, C*H*Me$_2$), 4.09 (1H, d, *J* = 7.9 Hz, C*H*OMe), 3.31 (3H, s, OC*H*$_3$), 1.54 (9H, s, C(C*H*$_3$)$_3$), 1.33 (6H, d, *J* = 6.1 Hz, CH(C*H*$_3$)$_2$) ppm; ^{13}C-NMR (100 MHz, DMSO-d$_6$): δ = 171.1 (q, CONH$_2$), 167.6 (q, *C*ONH), 165.3 (q, *C*ONH), 164.6 (q, CO$_2$*t*Bu), 164.3 (q, *C*ONH), 164.2 (q, *C*ONH), 152.1 (q, NH*C*ONH), 149.3 (q, C-Ar), 143.1 (q, C-Ar), 141.7 (q, C-Ar), 140.7 (q, C-Ar), 138.4 (q, C-Ar), 137.9

(q, C-Ar), 135.1 (q, C-Ar), 133.6 (t, *C*Hallyl), 130.1 (2C, t, C-Ar), 129.4 (2C, t, C-Ar), 129.1 (q, C-Ar), 128.3 (2C, t, C-Ar), 125.9 (q, C-Ar), 124.5 (t, C-Ar), 123.6 (2C, t, C-Ar), 122.9 (q, C-Ar), 120.1 (2C, t, C-Ar), 119.7 (2C, t, C-Ar), 118.8 (2C, t, C-Ar), 118.6 (2C, t, C-Ar), 117.9 (s, *C*H$_2$allyl), 113.7 (t, C-Ar), 80.3 (q, *C*(CH$_3$)$_3$), 80.1 (t, *C*HOMe), 75.9 (t, *C*H(CH$_3$)$_2$), 74.1 (s, OC*H*$_2$allyl), 57.7 (p, O*C*H$_3$), 55.6 (t, *C*HNH), 27.9 (3C, p, C(*C*H$_3$)$_3$), 21.9 (2C, p, CH(*C*H$_3$)$_2$) ppm; HRMS (ESI): *m/z* calc. for $C_{50}H_{52}N_8O_{13}Na$ [M + Na]$^+$: 995.3552; found 995.3553.

***tert*-Butyl-4-(2-(allyloxy)-4-(3-(4-((2*S*,3*R*)-4-amino-2-(4-(4-cyanobenzamido)-benzamido)-3-methoxy-4-oxobutanamido)phenyl)ureido)-3-isopropoxy-benzamido)benzoate (22):** Following the general procedure II acid **20** (34.5 mg, 129 µmol, 2.5 eq) and amine **18** (36.5 mg, 51.8 µmol, 1.0 eq) provided compound **22** as a colourless solid (26.6 mg, 27.9 µmol, 54%). Preparative HPLC (RP-18; run time 100 min; H$_2$O/MeCN = 70 : 30 → 0 : 100 in 80 min; t_r = 56 min). T_M = 239 °C (decomposition); $[\alpha]_D^{23}$ = +24.4° (*c* 1.2, MeOH); ^{1}H-NMR (600 MHz, DMSO-d$_6$): δ = 10.7 (1H, s, N*H*), 10.4 (1H, s, N*H*), 10.1 (1H, s, N*H*), 9.54 (1H, s, N*H*$_{\text{Urea}}$), 8.39 (1H, d, *J* = 8.1 Hz, N*H*CH), 8.31 (1H, s, N*H*$_{\text{Urea}}$), 8.13 – 8.10 (3H, m, Ar*H*), 8.04 (2H, m, Ar*H*), 7.94 – 7.81 (6H, m, Ar*H*), 7.81 (2H, m, Ar*H*), 7.60 (2H, m, Ar*H*), 7.53 (1H, s, CON*H*$_2$), 7.44 – 7.40 (4H, m, Ar*H*, CON*H*$_2$), 6.05 – 6.00 (1H, m, C*H*$_{\text{allyl}}$), 5.40 (1H, dq, *J* = 1.6, 17.1 Hz, C*H*$_{\text{allyl-}trans}$), 5.22 (1H, dq, *J* = 1.3, 10.4 Hz, C*H*$_{\text{allyl-}cis}$), 4.88 (1H, t, *J* = 8.1 Hz, C*H*NH), 4.63 – 4.59 (3H, m, OC*H*$_2$allyl, C*H*Me$_2$), 4.07 (1H, d, *J* = 8.1 Hz, C*H*OMe), 3.30 (3H, s, OC*H*$_3$), 1.54 (9H, s, C(C*H*$_3$)$_3$), 1.33 (6H, d, *J* = 6.1 Hz, CH(C*H*$_3$)$_2$) ppm; ^{13}C-NMR (125 MHz, DMSO-d$_6$): δ = 171.1 (q, CONH$_2$), 167.6 (q, *C*ONH), 165.3 (q, *C*ONH), 164.6 (q, CO$_2$*t*Bu), 164.5 (q, *C*ONH), 164.3 (q, *C*ONH), 152.0 (q, NH*C*ONH), 149.3 (q, C-Ar), 143.1 (q, C-Ar), 141.7 (q, C-Ar), 138.7 (q, C-Ar), 138.4 (q, C-Ar), 137.8 (q, C-Ar), 135.0 (q, C-Ar), 133.6 (t, *C*Hallyl), 132.5 (2C, t, C-Ar), 130.1 (2C, t, C-Ar), 129.0 (q, C-Ar), 128.6 (2C, t, C-Ar), 128.2 (2C, t, C-Ar), 125.9 (q, C-Ar), 124.5 (t, C-Ar), 123.0 (q, C-Ar), 120.1 (2C, t, C-Ar), 119.6 (2C, t, C-Ar), 118.8 (2C, t, C-Ar), 118.6 (2C, t, C-Ar), 118.3 (q, C-Ar), 117.9 (s, *C*H$_2$allyl), 117.3 (q, CN), 114.0 (q, C-Ar), 113.7 (t, C-Ar), 80.3 (q, *C*(CH$_3$)$_3$), 80.1 (t, *C*HOMe), 75.9 (t, *C*H(CH$_3$)$_2$), 74.1 (s, OC*H*$_2$allyl), 57.7 (p, O*C*H$_3$), 55.5 (t, *C*HNH), 27.9 (3C, p, C(*C*H$_3$)$_3$), 22.0 (2C, p, CH(*C*H$_3$)$_2$) ppm; HRMS (ESI): *m/z* calc. for $C_{51}H_{53}N_8O_{11}$ [M + H]$^+$: 953.3834; found 953.3840.

***tert*-Butyl-4-(4-(3-(4-((2*S*,3*R*)-4-amino-3-methoxy-2-(4-(4-nitrobenzamido)-benzamido)-4-oxobutanamido)phenyl)ureido)-2-hydroxy-3-isopropoxy-benzamido)benzoate (S6):** Following the general procedure VI compound **21** (25.0 mg, 25.7 µmol, 1.0 eq) provided phenol **S6** as a yellow amorphous solid (11.5 mg, 12.3 µmol, 48%). Preparative HPLC (RP-18; run time 80 min; H$_2$O/MeCN = 90 : 10 → 0 : 100 in 70 min; t_r = 61 min). $[\alpha]_D^{24}$ = +130.6° (*c* 0.9, MeCN); ^{1}H-NMR (400 MHz, DMSO-d$_6$): δ = 12.5 (1H, s, O*H*), 10.8 (1H, s, N*H*), 10.6 (1H, s, N*H*), 10.2 (1H, s, N*H*), 9.61 (1H, s, N*H*$_{\text{urea}}$), 8.39 – 8.37 (3H, m, Ar*H*, N*H*CH), 8.33 (1H, s, N*H*$_{\text{urea}}$), 8.21 (2H, m, Ar*H*), 7.92 – 7.79 (10H, m, Ar*H*), 7.60 (2H, m, Ar*H*), 7.53 (1H, s, CON*H*$_2$), 7.44 – 7.41 (3H, m, Ar*H*, CON*H*$_2$), 4.89 (1H, t, *J* = 8.0 Hz, C*H*NH), 4.63 (1H, hept, *J* = 6.1 Hz, C*H*Me$_2$), 4.07 (1H, d, *J* = 8.0 Hz, C*H*OMe), 3.31 (3H, s, OC*H*$_3$), 1.55 (9H, s, C(C*H*$_3$)$_3$), 1.30 (6H, d, *J* = 6.1 Hz, CH(C*H*$_3$)$_2$) ppm; ^{13}C-NMR (100 MHz, DMSO-d$_6$): δ = 171.1 (q, CONH$_2$), 169.0 (q, *C*ONH), 167.6 (q, *C*ONH), 165.4 (q, *C*ONH), 164.6 (q, CO$_2$*t*Bu), 164.2 (q, *C*ONH), 151.9 (q, NH*C*ONH), 149.3 (q, C-Ar), 142.1 (q, C-Ar), 141.7 (q, C-Ar), 140.3 (q, C-Ar), 139.0 (q, C-Ar), 135.0 (q, C-Ar), 133.6 (q, C-Ar), 132.9 (q, C-Ar), 129.9 (2C, t, C-Ar), 129.4 (2C, t, C-Ar), 129.1 (q, C-Ar), 128.3 (2C, t, C-Ar), 126.7 (q, C-Ar), 123.6 (2C, t, C-Ar), 123.0 (t, C-Ar), 120.7 (2C, t, C-Ar), 120.2 (2C, t, C-Ar), 119.7 (2C, t, C-Ar), 118.7 (2C, t, C-Ar), 109.3 (t, C-Ar), 80.5 (q, *C*(CH$_3$)$_3$), 80.1 (t, *C*HOMe), 74.2 (t, *C*H(CH$_3$)$_2$), 57.7 (p, O*C*H$_3$), 55.5 (t, *C*HNH), 27.9 (3C, p, C(*C*H$_3$)$_3$), 22.0 (2C, p, CH(*C*H$_3$)$_2$) ppm; HRMS (ESI): *m/z* calc. for $C_{47}H_{47}N_8O_{13}$ $[M - H]^-$: 931.3268; found 931.3262.

***tert*-Butyl-4-(4-(3-(4-((2*S*,3*R*)-4-amino-2-(4-(4-cyanobenzamido)benzamido)-3-methoxy-4-oxobutanamido)phenyl)ureido)-2-hydroxy-3-isopropoxybenz-amido)benzoate (S7):** Following the general procedure VI compound **22** (26.0 mg, 27.3 µmol, 1.0 eq) was used. Preparative HPLC (RP-18; run time 80 min; H$_2$O/MeCN = 70 : 30 → 0 : 100 in 70 min; t$_r$ = 26 min) $[\alpha]_D^{25}$ = +66.7° (*c* 0.8, MeOH); ^{1}H-NMR (600 MHz, DMSO-d$_6$): δ = 12.5 (1H, br. s, O*H*), 10.7 (1H, s, N*H*), 10.5 (1H, s, N*H*), 10.1 (1H, s, N*H*), 9.62 (1H, s, N*H*$_{\text{urea}}$), 8.39 (1H, d, *J* = 8.1 Hz, N*H*CH), 8.34 (1H, s, N*H*$_{\text{urea}}$), 8.13 (2H, m, Ar*H*), 8.04 (2H, m, Ar*H*), 7.93 – 7.87 (7H, m, Ar*H*), 7.84 (2H, m, Ar*H*), 7.81 (1H, d, *J* = 9.3 Hz, Ar*H*), 7.60 (2H, m, Ar*H*), 7.53 (1H, s, CON*H*$_2$), 7.44 – 7.42 (3H, m, Ar*H*, CON*H*$_2$), 4.89 (1H, t, *J* = 8.1 Hz, C*H*NH), 4.63 (1H, hept, *J* = 6.2 Hz, C*H*Me$_2$), 4.07 (1H, d, *J* = 8.1 Hz, C*H*OMe), 3.31 (3H, s, OC*H*$_3$), 1.55 (9H, s, C(C*H*$_3$)$_3$), 1.30 (6H, d, *J* = 6.2 Hz, CH(C*H*$_3$)$_2$) ppm; ^{13}C-NMR (125 MHz, DMSO-d$_6$): δ = 171.1 (q, CONH$_2$), 169.0 (q, *C*ONH), 167.6 (q, *C*ONH), 165.3 (q, *C*ONH), 164.6 (q, CO$_2$*t*Bu), 164.5 (q, *C*ONH), 154.4 (q, C-Ar), 151.9 (q, NH*C*ONH), 142.1 (q, C-Ar), 141.7 (q, C-Ar), 139.1 (q, C-Ar), 138.7 (q, C-Ar), 135.0 (q, C-Ar), 133.6 (q, C-Ar), 132.8 (q, C-Ar), 132.5 (2C, t, C-Ar), 129.9 (2C, t, C-Ar), 129.0 (q, C-Ar), 128.6 (2C, t, C-Ar), 128.2 (2C, t, C-Ar), 126.7 (q, C-Ar), 123.0 (t, C-Ar), 120.7 (2C, t, C-Ar), 120.1 (2C, t, C-Ar), 119.6 (2C, t, C-Ar), 118.6 (2C, t, C-Ar), 118.3 (q, CN), 114.0 (q, C-Ar), 109.2 (q, C-Ar), 108.1 (t, C-Ar), 80.5 (q, *C*(CH$_3$)$_3$), 80.1 (t, *C*HOMe), 74.2 (t, *C*H(CH$_3$)$_2$), 57.7 (p, O*C*H$_3$), 55.5 (t, *C*HNH), 27.8 (3C, p, C(*C*H$_3$)$_3$), 21.9 (2C, p, CH(*C*H$_3$)$_2$) ppm; HRMS (ESI): *m/z* calc. for $C_{48}H_{48}N_8O_{11}Na$ $[M + Na]^+$: 935.3340; found 935.3339.

4-(4-(3-(4-((2*S*,3*R*)-4-Amino-3-methoxy-2-(4-(4-nitrobenzamido)benzamido)-4-oxobutan-amido)phenyl)ureido)-2-hydroxy-3-isopropoxybenzamido)-benzoic acid (23): Following the general procedure V phenol **S6** (9.0 mg, 9.64 µmol, 1.0 eq) was stirred for 6 h. Acid **23** was obtained as a colourless amorphous solid (5.8 mg, 6.62 µmol, 69%). $[\alpha]_D^{24}$ = +38.7° (*c* 0.6, DMSO); ^{1}H-NMR (500

MHz, DMSO-d_6 +2 µL DCO_2D): δ = 10.8 (1H, s, N*H*), 10.6 (1H, br. s, N*H*), 10.2 (1H, s, N*H*), 9.61 (1H, s, NH_{Urea}), 8.40 – 8.37 (3H, m, Ar*H*, N*H*CH), 8.33 (1H, s, NH_{Urea}), 8.20 (2H, m, Ar*H*), 7.96 (2H, m, Ar*H*), 7.92 – 7.84 (5H, m, Ar*H*), 7.84 (2H, m, Ar*H*), 7.80 (1H, d, *J* = 8.5 Hz, Ar*H*), 7.60 (2H, m, Ar*H*), 7.49 (2H, d, *J* = 46.1 Hz, CONH_2), 7.42 (2H, m, Ar*H*), 4.89 (1H, t, *J* = 8.5 Hz, C*H*NH), 4.63 (1H, hept, *J* = 6.0 Hz, C*H*Me_2), 4.07 (1H, d, *J* = 7.3 Hz, C*H*OMe), 3.30 (3H, s, OCH_3), 1.30 (6H, d, *J* = 6.0 Hz, CH(CH_3)$_2$) ppm; ^{13}C-NMR (125 MHz, DMSO-d_6 +2 µL DCO_2D): δ = 171.1 (q, $CONH_2$), 169.0 (q, *C*ONH), 167.6 (q, *C*ONH), 167.0 (q, *C*O_2H), 165.4 (q, *C*ONH), 164.3 (q, *C*ONH), 154.5 (q, C-Ar), 152.0 (q, NH*C*ONH), 149.3 (q, C-Ar), 142.1 (q, C-Ar), 141.7 (q, C-Ar), 140.4 (q, C-Ar), 139.1 (q, C-Ar), 135.0 (q, C-Ar), 133.7 (q, C-Ar), 132.9 (q, C-Ar), 130.2 (2C, t, C-Ar), 129.4 (2C, t, C-Ar), 129.2 (q, C-Ar), 128.3 (2C, t, C-Ar), 126.2 (q, C-Ar), 123.6 (2C, t, C-Ar), 123.1 (t, C-Ar), 120.8 (2C, t, C-Ar), 120.2 (2C, t, C-Ar), 119.7 (2C, t, C-Ar), 118.7 (2C, t, C-Ar), 109.3 (q, C-Ar), 108.2 (t, C-Ar), 80.2 (t, *C*HOMe), 74.2 (t, *C*H(CH_3)$_2$), 57.7 (p, O*C*H_3), 55.6 (t, *C*HNH), 22.0 (2C, p, CH(*C*H_3)$_2$) ppm; HRMS (ESI): *m/z* calc. for $C_{43}H_{40}N_8O_{13}Na$ $[M + Na]^+$: 899.2613; found 899.2613.

4-(4-(3-(4-((2*S*,3*R*)-4-Amino-2-(4-(4-cyanobenzamido)benzamido)-3-methoxy-4-oxo-butanamido)phenyl)ureido)-2-hydroxy-3-isopropoxybenz-amido)benzoic acid (24): Following the general procedure V phenol **S7** (14.0 mg, 15.3 µmol, 1.0 eq) was stirred for 30 min. Acid **24** was obtained as a yellow amorphous solid (13.0 mg, 15.2 µmol, 99%). $[\alpha]_D^{23}$ = +37.2° (*c* 0.3, DMSO); ^{1}H-NMR (500 MHz, DMSO-d_6 +2 µL DCO_2D): δ = 12.5 (1H, s, O*H*), 10.7 (1H, s, N*H*), 10.5 (1H, s, N*H*), 10.1 (1H, s, N*H*), 9.61 (1H, s, NH_{Urea}), 8.38 (1H, d, *J* = 8.1 Hz, N*H*CH), 8.34 (1H, s, NH_{Urea}), 8.13 (2H, m, Ar*H*), 8.04 (2H, m, Ar*H*), 7.96 (2H, m, Ar*H*), 7.91 – 7.83 (7H, m, Ar*H*), 7.81 (1H, d, *J* = 9.2 Hz, Ar*H*), 7.60 (2H, m, Ar*H*), 7.52 – 7.41 (4H, m, Ar*H*, CONH_2), 4.88 (1H, t, *J* = 8.1 Hz, C*H*NH), 4.62 (1H, hept, *J* = 6.1 Hz, C*H*Me_2), 4.06 (1H, d, *J* = 8.1 Hz, C*H*OMe), 3.30 (3H, s, OCH_3), 1.30 (6H, d, *J* = 6.1 Hz, CH(CH_3)$_2$) ppm; ^{13}C-NMR (125 MHz, DMSO-d_6 +2 µL DCO_2D): δ = 171.1 (q, $CONH_2$), 169.0 (q, *C*ONH), 167.6 (q, *C*ONH), 166.9 (q, *C*O_2H), 165.3 (q, *C*ONH), 164.5 (q, *C*ONH), 154.3 (q, C-Ar), 151.9 (q, NH*C*ONH), 142.0 (q, C-Ar), 141.7 (q, C-Ar), 139.1 (q, C-Ar), 138.7 (q, C-Ar), 134.9 (q, C-Ar), 133.7 (q, C-Ar), 132.5 (2C, t, C-Ar), 130.2 (2C, t, C-Ar), 129.0 (q, C-Ar), 128.6 (2C, t, C-Ar), 128.3 (2C, t, C-Ar), 126.1 (q, C-Ar), 123.0 (t, C-Ar), 120.8 (2C, t, C-Ar), 120.6 (q, C-Ar), 120.1 (2C, t, C-Ar), 119.6 (2C, t, C-Ar), 118.7 (2C, t, C-Ar), 118.3 (q, CN), 114.1 (q, C-Ar), 109.2 (q, C-Ar), 108.2 (t, C-Ar), 80.1 (t, *C*HOMe), 74.3 (t, *C*H(CH_3)$_2$), 57.7 (p, O*C*H_3), 55.5 (t, *C*HNH), 21.9 (2C, p, CH(*C*H_3)$_2$) ppm; HRMS (ESI): *m/z* calc. for $C_{44}H_{40}N_8O_{11}Na$ $[M + Na]^+$: 879.2714; found 879.2714.

***tert*-Butyl-4-(2-(allyloxy)-4-(3-(4-((2*S*,3*R*)-4-amino-2-(4-(3-(4-cyanophenyl)-ureido)-benz-amido)-3-methoxy-4-oxobutanamido)phenyl)ureido)-3-iso-propoxybenzamido)-benzoate (26):** Following the general procedure II acid **8** (26.0 mg, 92.2 µmol, 2.5 eq) and amine **18** (26.0 mg, 36.9 µmol, 1.0 eq) provided compound **26** as a colourless solid (27.0 mg, 27.9 µmol, 76%). Preparative HPLC (RP-18; run time 100 min; H_2O/MeCN = 70 : 30 → 0 : 100 in 80 min; t_r = 56 min). T_M = 203 °C (decomposition); $[\alpha]_D^{24}$ = +40.0° (*c* 0.8, MeOH); ^{1}H-NMR (500 MHz, DMSO-d_6): δ = 10.4 (1H, s, N*H*), 10.1 (1H, s, N*H*), 9.52 (1H, s, NH_{urea}), 9.43 (1H, s, NH_{urea}), 9.30 (1H, s, NH_{urea}), 8.31 – 8.30 (2H, m, NH_{urea}, N*H*CH), 8.11 (1H, d, *J* = 8.8 Hz, Ar*H*), 7.88 (2H, m, Ar*H*), 7.82 – 7.80 (4H, m, Ar*H*), 7.74 (2H, m, Ar*H*), 7.65 (2H, m, Ar*H*), 7.60 – 7.57 (4H, m, Ar*H*), 7.51 (1H, s, CONH_2), 7.43 – 7.40 (4H, m, Ar*H*, CONH_2), 6.07 – 5.99 (1H, m, CH_{allyl}), 5.40 (1H, dd, *J* = 1.7, 17.2 Hz, C$H_{allyl\text{-}trans}$), 5.22 (1H, dd, *J* = 1.5, 10.5 Hz, C$H_{allyl\text{-}cis}$), 4.86 (1H, t, *J* = 8.1 Hz, C*H*NH), 4.64 – 4.59 (3H, m, OCH_2allyl, C*H*Me_2), 4.06 (1H, d, *J* = 8.1 Hz, C*H*OMe), 3.30 (3H, s, OCH_3), 1.54 (9H, s, C(CH_3)$_3$), 1.32 (6H, d, *J* = 6.2 Hz, CH(CH_3)$_2$) ppm; ^{13}C-NMR (125 MHz, DMSO-d_6): δ = 171.1 (q, $CONH_2$), 167.7 (q, *C*ONH), 165.4 (q, *C*ONH), 164.6 (q, *C*ONH), 164.3 (q, CO_2*t*Bu), 152.0 (q, NH*C*ONH), 152.0 (q, NH*C*ONH), 149.3 (q, C-Ar), 144.0 (q, C-Ar), 143.1 (q, C-Ar), 142.4 (q, C-Ar), 138.4 (q, C-Ar), 137.8 (q, C-Ar), 135.0 (q, C-Ar), 133.6 (q, C-Ar), 133.6 (t, *C*Hallyl), 133.3 (2C, t, C-Ar), 130.1 (2C, t, C-Ar), 128.5 (2C, t, C-Ar), 127.2 (q, C-Ar), 125.9 (q, C-Ar), 124.4 (t, C-Ar), 123.0 (q, C-Ar), 120.1 (2C, t, C-Ar), 119.3 (q, CN), 118.8 (2C, t, C-Ar), 118.6 (2C, t, C-Ar), 118.2 (2C, t, C-Ar), 117.9 (s, *C*H_2allyl), 117.5 (2C, t, C-Ar), 113.7 (t, C-Ar), 103.5 (q, C-Ar), 80.3 (q, *C*(CH_3)$_3$), 80.1 (t, *C*HOMe), 75.9 (t, *C*H(CH_3)$_2$), 74.1 (s, OCH_2allyl), 57.7 (p, O*C*H_3), 55.5 (t, *C*HNH), 27.9 (3C, p, C(*C*H_3)$_3$), 21.9 (2C, p, CH(*C*H_3)$_2$) ppm; HRMS (ESI): *m/z* calc. for $C_{51}H_{53}N_9O_{11}Na$ $[M + Na]^+$: 990.3762; found 990.3769.

***tert*-Butyl-4-(4-(3-(4-((2*S*,3*R*)-4-amino-2-(4-(3-(4-cyanophenyl)ureido)-benz-amido)-3-methoxy-4-oxobutanamido)phenyl)ureido)-2-hydroxy-3-iso-propoxybenzamido)benzoate (S8):** Following the general procedure VI compound **26** (26.0 mg, 26.8 µmol, 1.0 eq) was used. Preparative HPLC (RP-18; run time 80 min; H_2O/MeCN = 70 : 30 → 0 : 100 in 70 min; t_r = 61 min) provided phenol **S8** as a colourless amorphous solid (9.3 mg, 10.0 µmol, 37%). $[\alpha]_D^{22}$ = +43.6° (*c* 0.9, MeOH); ^{1}H-NMR (600 MHz, DMSO-d_6): δ = 12.5 (1H, br. s, O*H*), 10.6 (1H, br. s, N*H*), 10.1 (1H, s, N*H*), 9.61 (1H, s, NH_{urea}), 9.50 (1H, s, NH_{urea}), 9.36 (1H, s, NH_{urea}), 9.32 (1H, s, NH_{urea}), 8.30 (1H, d, *J* = 8.1 Hz, N*H*CH), 7.91 (2H, m, Ar*H*), 7.88 – 7.79 (6H, m, Ar*H*), 7.73 (2H, m, Ar*H*), 7.66 (2H, m, Ar*H*), 7.60 – 7.56 (4H, m, Ar*H*), 7.52 – 7.38 (4H, m, Ar*H*, CONH_2), 4.86 (1H, t, *J* = 8.1 Hz, C*H*NH), 4.63 (1H, hept, *J* = 6.2 Hz, C*H*Me_2), 4.06 (1H, d, *J* = 8.1 Hz, C*H*OMe), 3.30 (3H, s, OCH_3), 1.55 (9H, s, C(CH_3)$_3$), 1.30 (6H, d, *J* = 6.2 Hz, CH(CH_3)$_2$) ppm; ^{13}C-NMR (125 MHz, DMSO-d_6): δ = 171.1 (q, $CONH_2$), 169.0 (q, *C*ONH), 167.7 (q, *C*ONH), 165.4 (q, *C*ONH), 164.6 (q, CO_2*t*Bu), 154.6 (q, C-Ar), 152.0 (q, NH*C*ONH), 151.9 (q, NH*C*ONH), 144.0 (q, C-Ar), 142.4 (q, C-Ar), 142.2 (q, C-Ar), 139.0 (q, C-Ar), 135.0 (q, C-Ar), 133.6 (q, C-Ar), 133.3 (2C, t, C-Ar), 132.9 (q, C-Ar), 129.9 (2C, t, C-Ar), 128.5 (2C, t, C-Ar), 127.2 (q, C-Ar), 126.6 (q, C-Ar), 123.0 (t, C-Ar), 120.6 (2C, t, C-Ar), 120.1 (2C, t, C-Ar), 119.3 (q, CN), 118.6 (2C, t, C-Ar), 118.2 (2C, t, C-Ar), 117.5 (2C, t, C-Ar), 109.3 (q, C-Ar), 107.9 (t, C-Ar), 103.5 (q, C-Ar), 80.4 (q, *C*(CH_3)$_3$), 80.1 (t, *C*HOMe), 74.1 (t, *C*H(CH_3)$_2$), 57.7 (p, O*C*H_3), 55.5 (t, *C*HNH), 27.8 (3C, p, C(CH_3)$_3$), 21.9 (2C, p, CH(CH_3)$_2$) ppm; HRMS (ESI): *m/z* calc. for $C_{48}H_{49}N_9O_{11}Na$ [M + Na]$^+$: 950.3449; found 950.3447.

4-(4-(3-(4-((2*S*,3*R*)-4-Amino-2-(4-(3-(4-cyanophenyl)ureido)benzamido)-3-methoxy-4-oxobutanamido)phenyl)ureido)-2-hydroxy-3-isopropoxybenz-amido)benzoic acid (27): Following the general procedure V phenol **S8** (9.0 mg, 9.70 µmol, 1.0 eq) was stirred for 30 min. Acid **27** was obtained as a colourless amorphous solid (8.4 mg, 9.70 µmol, 99%). $[\alpha]_D^{23}$ = +12.7° (*c* 1.1, MeOH); ^{1}H-NMR (600 MHz, DMSO-d$_6$ +2 µL DCO$_2$D): δ = 12.5 (1H, br. s, O*H*), 10.5 (1H, br. s, N*H*), 10.1 (1H, s, N*H*), 9.61 (1H, s, NH_{Urea}), 9.37 (1H, s, NH_{Urea}), 9.25 (1H, s, NH_{Urea}), 8.34 (1H, s, NH_{Urea}),8.30 (1H, d, *J* = 8.1 Hz, N*H*CH), 7.96 (2H, m, Ar*H*), 7.89 (1H, d, *J* = 9.0 Hz, Ar*H*), 7.85 – 7.80 (5H, m, Ar*H*), 7.74 (2H, m, Ar*H*), 7.65 (2H, m, Ar*H*), 7.60 – 7.55 (4H, m, Ar*H*), 7.52 – 7.41 (4H, m, Ar*H*, CONH_2), 4.86 (1H, t, *J* = 8.1 Hz, C*H*NH), 4.62 (1H, hept, *J* = 6.2 Hz, C*H*Me$_2$), 4.06 (1H, d, *J* = 8.1 Hz, C*H*OMe), 3.30 (3H, s, OCH_3), 1.30 (6H, d, *J* = 6.2 Hz, CH(CH_3)$_2$) ppm; ^{13}C-NMR (125 MHz, DMSO-d$_6$ +2 µL DCO$_2$D): δ = 171.1 (q, CONH$_2$), 169.0 (q, *C*ONH), 167.7 (q, *C*ONH), 166.9 (q, *C*O$_2$H), 165.4 (q, *C*ONH), 154.3 (q, C-Ar), 152.0 (q, NH*C*ONH), 151.9 (q, NH*C*ONH), 144.0 (q, C-Ar), 142.3 (q, C-Ar), 142.0 (q, C-Ar), 139.1 (q, C-Ar), 134.9 (q, C-Ar), 133.7 (q, C-Ar), 133.3 (2C, t, C-Ar), 132.8 (q, C-Ar), 130.2 (2C, t, C-Ar), 128.5 (2C, t, C-Ar), 127.3 (q, C-Ar), 126.2 (q, C-Ar), 123.0 (t, C-Ar), 120.8 (2C, t, C-Ar), 120.1 (2C, t, C-Ar), 119.3 (q, CN), 118.7 (2C, t, C-Ar), 118.2 (2C, t, C-Ar), 117.5 (2C, t, C-Ar), 109.2 (q, C-Ar), 108.2 (t, C-Ar), 103.5 (q, C-Ar), 80.1 (t, *C*HOMe), 74.3 (t, *C*H(CH$_3$)$_2$), 57.7 (p, O*C*H$_3$), 55.5 (t, *C*HNH), 21.9 (2C, p, CH(*C*H$_3$)$_2$) ppm; HRMS (ESI): *m/z* calc. for $C_{44}H_{42}N_9O_{11}$ [M + H]$^+$: 872.3004; found 872.3004.

***tert*-Butyl-4-(2-(allyloxy)-4-azido-3-isopropoxybenzamido)benzoate (28):** *tert*-Butylnitrite (1.13 mL, 9.50 mmol, 1.5 eq) and trimethylsilyl azide (1.26 mL, 9.50 mmol, 1.5 eq) were added dropwise to a solution of amine **9** (2.70 g, 6.34 mmol, 1.0 eq) in acetonitrile (127 mL) at 0 °C. After 2 h at rt all volatiles were removed *in vacuo*. Column chromatography (PE/EtOAc = 20:1 → 10:1) yielded azide **28** (2.86 g, 6.34 mmol, quant.) as a yellow solid. T_M = 78 °C; ^{1}H-NMR (400 MHz, CDCl$_3$): δ = 10.2 (1H, s, H-8), 8.00 – 7.96 (3H, m, H-5, H-5‘, H-11), 7.71 (2H, m, H-6, H-6‘), 6.95 (1H, d, *J* = 8.7 Hz, H-12), 6.17 – 6.08 (1H, m, H-19), 5.48 (1H, dq, *J* = 1.4, 17.2 Hz, H-20$_{trans}$), 5.40 (1H, dq, *J* = 1.2, 10.3 Hz, H-20$_{cis}$), 4.73 (2H, dt, *J* = 1.2, 6.0 Hz, H-18), 4.60 (1H, hept, *J* = 6.2 Hz, H-16), 1.60 (9H, s, H-1), 1.36 (6H, d, *J* = 6.2 Hz, H-17) ppm; ^{13}C-NMR (100 MHz, CDCl$_3$): δ = 165.5 (q, C-3), 162.5 (q, C-9), 151.4 (q, C-15), 143.1 (q, C-14), 142.2 (q, C-7), 139.2 (q, C-13), 132.3 (t, C-19), 130.8 (2C, t, C-5, C-5’), 127.6 (q, C-4), 127.1 (t, C-11), 123.3 (q, C-10), 120.3 (s, C-20), 119.2 (2C, t, C-6, C-6’), 116.4 (t, C-12), 81.0 (q, C-2), 76.9 (t, C-16), 75.3 (s, C-18), 28.4 (3C, p, C-1), 22.3 (2C, p, C-17) ppm; HRMS (ESI): *m/z* calc. for $C_{24}H_{28}N_4O_5Na$ $[M + Na]^+$: 475.1957; found 475.1959; IR (neat): $\tilde{\nu}$ = 3318, 2980, 2118, 1708, 1672, 1589, 1528, 1252, 1101, 978, 770 cm^{-1}.

***tert*-Butyl-4-(2-(allyloxy)-4-(4-(4-aminophenyl)-1*H*-1,2,3-triazol-1-yl)-3-iso-propoxybenzamido)benzoate (30):** Azide **28** (2.98 g, 6.58 mmol, 1.0 eq) and 4-ethynylaniline (**29**, 771 mg, 6.56 mmol, 1.0 eq) were dissolved in a mixture of THF (55 mL) and DMSO (110 mL). Sodium ascorbate (782 mg, 3.95 mmol, 0.6 eq) was dissolved in water (5.5 mL) and added to the reaction mixture. After addition of tris[(1-benzyl-1*H*-1,2,3-triazol-4-yl)methyl]amine (1.40 g, 2.63 mmol, 0.4 eq) and $CuSO_4$ (105 mg, 0.66 mmol, 0.1 eq), the reaction was stirred at rt for 2 h. Half of the solvent was removed under reduced pressure. The residue was extracted with EtOAc (3x). The combined organic phases were washed with NH_4Cl-solution, water and brine, dried over Na_2SO_4, filtered and concentrated *in vacuo*. Column chromatography (PE/EtOAc = 20:1) yielded the desired product **30** (3.56 g, 6.25 mmol, 95%) as yellow solid. T_M = 75 °C; ^{1}H-NMR (400 MHz, $CDCl_3$): δ = 10.2 (1H, s, H-8), 8.36 (1H, s, H-21), 8.16 (1H, d, *J* = 8.7 Hz, H-11), 8.00 (2H, m, H-5, H-5‘), 7.80 (1H, d, *J* = 8.7 Hz, H-12), 7.76 – 7.72 (4H, m, H-6, H-6‘, H-24, H-24‘), 6.38 (2H, m, H-25, H-25‘), 6.22 – 6.12 (1H, m, H-19), 5.53 (1H, dq, *J* = 1.2, 17.1 Hz, H-20_{trans}), 5.44 (1H, dq, *J* = 1.0, 10.4 Hz, H-20_{cis}), 4.83 (2H, d, *J* = 6.0 Hz, H-18), 4.36 (1H, hept, *J* = 6.2 Hz, H-16), 1.60 (9H, s, H-1), 1.12 (6H, d, *J* = 6.2 Hz, H-17) ppm; ^{13}C-NMR (100 MHz, $CDCl_3$): δ = 165.4 (q, C-3), 162.0 (q, C-9), 151.4 (q, C-15), 148.1 (q, C-22), 146.0 (q, C-26), 142.9 (q, C-14), 141.9 (q, C-7), 135.8 (q, C-13), 132.1 (t, C-19), 130.8 (2C, t, C-5, C-5’), 127.8 (q, C-4/C-10), 127.3 (q, C-4/C-10), 127.3 (t, C-11), 127.2 (2C, t, C-24, C-24’), 121.3 (q, C-23), 120.8 (t, C-12), 120.6 (s, C-20), 120.3 (t, C-21), 119.3 (2C, t, C-6, C-6’), 115.9 (2C, t, C-25, C-25’), 81.0 (q, C-2), 77.9 (t, C-16), 75.6 (s, C-18), 28.4 (3C, p, C-1), 22.3 (2C, p, C-17) ppm; HRMS (ESI): *m/z* calc. for $C_{32}H_{36}N_5O_5$ $[M + H]^+$: 570.2716; found 570.2714.

***tert*-Butyl-4-(2-(allyloxy)-4-(4-(4-((2*S*,3*R*)-4-amino-2-((*tert*-butoxycarbonyl)-amino)-3-methoxy-4-oxobutanamido)phenyl)-1H-1,2,3-triazol-1-yl)-3-iso-propoxybenzamido)-benzoate (S9):** Following the general procedure III using amine **30** (2.83 g, 4.97 mmol, 1.0 eq) and acid **13** (2.22 g, 8.45 mmol, 1.7 eq) compound **S9** (2.72 g, 3.34 mmol, 67%) was obtained as a yellow amorphous solid. $[\alpha]_D^{24}$ = -3.9° (*c* 2.0, MeOH); ^{1}H-NMR (400 MHz, DMSO-d_6): δ = 10.7 (1H, s, N*H*), 10.2 (1H, s, N*H*), 8.89 (1H, s, CH_{Triazol}), 7.93 – 7.91 (4H, m, Ar*H*), 7.84 (2H, m, Ar*H*), 7.76 (2H, m, Ar*H*), 7.61 (1H, d, *J* = 8.3 Hz, Ar*H*), 7.54 (1H, d, *J* = 8.3 Hz, Ar*H*), 7.43 (2H, d, *J* = 17.8 Hz, NH_2), 6.76 (1H, d, *J* = 8.4 Hz, N*H*Boc), 6.07 – 5.97 (1H, m, CH_{allyl}), 5.38 (1H, dq, *J* = 1.6, 17.3 Hz, C$H_{\text{allyl-}trans}$), 5.20 (1H, dd, *J* = 1.4, 9.6 Hz, C$H_{\text{allyl-}cis}$), 4.69 (2H, d, *J* = 5.5 Hz, OCH_2allyl), 4.40 (1H, t, *J* = 7.5 Hz, C*H*NH), 4.28 (1H, hept, *J* = 6.1 Hz, C*H*Me$_2$), 3.86 (1H, d, *J* = 7.4 Hz, C*H*OMe), 3.26 (3H, s, OCH_3), 1.55 (9H, s, C(CH_3)$_3$), 1.39 (9H, s, C(CH_3)$_3$), 1.01 (6H, d, *J* = 6.1 Hz, CH(CH_3)$_2$) ppm; ^{13}C-NMR (100 MHz, DMSO-d_6): δ = 170.7

(q, CONH$_2$), 168.4 (q, *C*ONH), 168.1 (q, CO$_2$*t*Bu), 164.6 (q, *C*ONH), 164.3 (q, CO$_2$*t*Bu), 150.3 (q, *C*-Ar), 146.3 (q, C-Ar$_{\mathit{Triazol}}$), 143.9 (q, C-Ar), 142.8 (q, C-Ar), 138.9 (q, C-Ar), 133.6 (q, C-Ar), 133.5 (t, *C*Hallyl), 132.9 (q, C-Ar), 130.1 (2C, t, C-Ar), 126.3 (q, C-Ar), 125.8 (2C, t, C-Ar), 125.3 (q, C-Ar), 123.9 (t, C-Ar), 122.8 (t, C-H$_{\mathit{Triazol}}$), 120.9 (t, C-Ar), 119.8 (2C, t, C-Ar), 118.9 (2C, t, C-Ar), 118.0 (s, *C*H$_2$allyl), 80.4 (t, *C*HOMe), 80.2 (q, *C*(CH$_3$)$_3$), 78.6 (q, *C*(CH$_3$)$_3$), 76.8 (t, *C*H(CH$_3$)$_2$), 74.6 (s, OC*H*$_2$allyl), 57.6 (p, O*C*H$_3$), 56.5 (t, *C*HNH), 28.1 (3C, p, C(*C*H$_3$)$_3$), 27.8 (3C, p, C(*C*H$_3$)$_3$), 21.8 (2C, p, CH(*C*H$_3$)$_2$) ppm; HRMS (ESI): *m/z* calc. for $C_{42}H_{51}N_7O_{10}Na$ [M + Na]$^+$: 836.33595; found 836.3596.

***tert*-Butyl-4-(2-(allyloxy)-4-(4-(4-((2*S*,3*R*)-2,4-diamino-3-methoxy-4-oxobutanamido)-phenyl)-1H-1,2,3-triazol-1-yl)-3-isopropoxybenzamido)-benzoate (31):** Following the general procedure IV compound **S9** (420 mg, 0.52 mmol, 1.0 eq) was used. Preparative HPLC (RP-18; run time 100 min; H$_2$O/MeCN = 80 : 20 → 0 : 100 in 80 min; t$_r$ = 31 min) provided amine **31** as a yellow amorphous solid (131 mg, 0.18 mmol, 36%). **$[\alpha]_D^{24}$** = +27.7° (*c* 1.1, MeOH); ^{1}H-NMR (400 MHz, DMSO-d$_6$): δ = 10.7 (1H, s, N*H*), 8.93 (1H, s, C*H*$_{\text{Triazol}}$), 8.14 – 7.73 (10H, m, N*H*, Ar*H*), 7.66 – 7.53 (3H, m, Ar*H*, N*H*$_2$), 6.07 – 5.99 (1H, m, C*H*$_{\text{allyl}}$), 5.39 (1H, d, *J* = 17.2 Hz, C*H*$_{\text{allyl-}\mathit{trans}}$), 5.20 (1H, d, *J* = 10.3 Hz, C*H*$_{\text{allyl-}\mathit{cis}}$), 4.69 (2H, d, *J* = 5.5 Hz, OC*H*$_2$allyl), 4.31 – 4.13 (3H, m, C*H*NH$_2$, C*H*OMe, C*H*Me$_2$), 3.39 (3H, s, OC*H*$_3$), 1.55 (9H, s, C(C*H*$_3$)$_3$), 1.00 (6H, d, *J* = 6.1 Hz, CH(C*H*$_3$)$_2$) ppm; ^{13}C-NMR (100 MHz, DMSO-d$_6$): δ = 170.0 (q, CONH$_2$), 164.6 (q, *C*ONH), 164.3 (q, *C*ONH), 163.0 (q, CO$_2$*t*Bu), 150.3 (q, *C*-Ar), 146.2 (q, C-Ar$_{\mathit{Triazol}}$), 143.9 (q, C-Ar), 142.8 (q, C-Ar), 138.2 (q, C-Ar), 133.6 (q, C-Ar), 133.5 (t, *C*Hallyl), 133.0 (q, C-Ar), 130.1 (2C, t, C-Ar), 126.3 (q, C-Ar), 126.0 (2C, t, C-Ar), 125.9 (q, C-Ar), 123.9 (t, C-Ar), 122.9 (t, C-H$_{\mathit{Triazol}}$), 120.9 (t, C-Ar), 120.0 (2C, t, C-Ar), 118.9 (2C, t, C-Ar), 118.0 (s, *C*H$_2$allyl), 80.4 (q, *C*(CH$_3$)$_3$), 78.7 (t, *C*HOMe), 76.8 (t, *C*H(CH$_3$)$_2$), 74.6 (s, OC*H*$_2$allyl), 58.6 (p, O*C*H$_3$), 54.3 (t, *C*HNH$_2$), 27.9 (3C, p, C(*C*H$_3$)$_3$), 21.9 (2C, p, CH(*C*H$_3$)$_2$) ppm; HRMS (ESI): *m/z* calc. for $C_{37}H_{44}N_7O_8$ [M + H]$^+$: 714.3251; found 714.3250.

4-(4-Fluorobenzamido)-benzoic acid (39): 4-Fluorbenzoyl chloride (**33**, 3.5 mL, 29.2 mmol, 4.0 eq) was dropped to a solution of 4-aminobenzoic acid (**7**, 1.0 g, 7.29 mmol, 1.0 eq) and Na$_2$CO$_3$ (1.55 g, 14.6 mmol, 2.0 eq) in water/THF (47 mL, 1:1). After stirring vigorously at rt for 3 h, the filtrate was filtered and washed with hot water (1.5 L). Acid **39** (1.85 g, 7.15 mmol, 98%) was obtained after drying at high vacuum as colourless solid. T$_M$ = 301 °C; ^{1}H-NMR (400 MHz, DMSO-d$_6$): δ = 12.8 (1H, s, CO$_2$*H*), 10.5 (1H, s, H-6), 8.07 – 8.03 (2H, m, H-9, H-9‘), 7.95 – 7.89 (4H, m, H-3, H-3‘, H-4, H-4‘), 7.41 – 7.36 (2H, m, H-10, H-10‘) ppm; ^{13}C-NMR (100 MHz, DMSO-d$_6$): δ = 166.9 (q, C-1), 164.8 (q, C-7), 164.2 (d, *J* = 249.4 Hz, C-11), 143.2 (q, C-5), 131.1 (d, *J* = 2.9 Hz, C-8), 130.6 (d, *J* = 9.2 Hz, C-9, C-9‘), 130.2 (2C, t, C-3, C-3‘), 125.5 (q, C-2), 119.5 (2C, t, C-4, C-4‘), 115.4 (d, *J* = 21.8 Hz, C-10, C-10‘) ppm; HRMS (ESI): *m/z* calc. for $C_{14}H_9NO_3F$ [M – H]$^-$: 258.0566; found 258.0564.

4-(5-Cyanopicolinamido)-benzoic acid (40): Following the general procedure I using amine **32** (32.6 mg, 0.17 mmol) and acid **34** (50 mg, 0.34 mmol) intermediate was isolated and used directly for *t*Bu-saponification following to general procedure V**.** Acid **40** (33.7 mg, 0.13 mmol, 75%) was obtained as a beige solid.T_M = 354 °C; ^{1}H-NMR (400 MHz, DMSO-d_6): δ = 12.8 (1H, s, CO_2*H*), 11.1 (1H, s, H-6), 9.22 (1H, d, *J* = 1.7 Hz, H-12), 8.60 (1H, dd, *J* = 1.7, 8.1 Hz, H-10), 8.31 (1H, d, *J* = 8.1 Hz, H-9), 8.06 (2H, m, H-4, H-4‘), 7.95 (2H, m, H-3, H-3‘) ppm; ^{13}C-NMR (100 MHz, DMSO-d_6): δ = 166.9 (q, C-1), 161.8 (q, C-7), 152.3 (q, C-8), 151.5 (t, C-10), 142.3 (t, C-12), 142.1 (q, C-5), 130.2 (2C, t, C-3, C-3‘), 126.2 (q, C-2), 122.6 (t, C-9), 120.0 (2C, t, C-4, C-4‘), 116.6 (q, CN), 111.8 (q, C-11) ppm; HRMS (ESI): *m/z* calc. for $C_{14}H_8N_3O_3$ $[M-H]^-$: 266.0566; found 266.0564.

4-(4-Cyano-3-fluorobenzamido)-benzoic acid (41): Following the general procedure I using amine **32** (58.5 mg, 0.30 mmol) and acid **35** (100 mg, 0.61 mmol) intermediate was isolated and used directly for *t*Bu-saponification following to general procedure V. Acid **41** (57.0 mg, 0.20 mmol, 66%) was obtained as a colourless solid.T_M = 298 °C; ^{1}H-NMR (400 MHz, DMSO-d_6): δ = 12.8 (1H, s, CO_2*H*), 10.8 (1H, s, H-6), 8.16 – 8.13 (1H, m, H-9), 8.08 – 8.05 (1H, m, H-13), 7.98 – 7.95 (3H, m, H-4, H-4‘, H-10), 7.91 – 7.88 (2H, m, H-3, H-3‘) ppm; ^{13}C-NMR (100 MHz, DMSO-d_6): δ = 166.8 (q, C-1), 163.5 (q, C-7), 162.1 (d, *J* = 238.1 Hz, C-12), 142.6 (q, C-5), 141.5 (d, *J* = 7.3 Hz, C-8), 134.3 (t, C-9), 130.3 (2C, t, C-3, C-3‘), 126.1 (q, C-2), 124.7 (d, *J* = 3.5 Hz, C-10), 119.7 (2C, t, C-4, C-4‘), 115.8 (d, *J* = 21.4 Hz, C-13), 113.6 (q, CN), 102.9 (d, *J* = 15.3 Hz, C-11) ppm; HRMS (ESI): *m/z* calc. for $C_{15}H_8N_2O_3F$ $[M-H]^-$: 283.0519; found 283.0516.

4-(Isonicotinamido)-benzoic acid (42): Following to general procedure I using amine **32** (78.4 mg, 0.41 mmol) and acid **36** (100 mg, 0.81 mmol) the intermediate was isolated and directly used for *t*Bu-saponification following the general procedure V. Acid **42** (92.3 mg, 0.38 mmol, 94%) was obtained as a beige solid. T_M = 381 °C; ^{1}H-NMR (400 MHz, DMSO-d_6): δ = 12.8 (1H, s, CO_2*H*), 10.8 (1H, s, H-6), 8.82 (2H, m, H-10, H-10‘), 7.97 (2H, m, H-4, H-4‘), 7.92 – 7.91 (2H, m, H-3, H-3‘), 7.90 – 7.89 (2H, m, H-9, H-9‘) ppm; ^{13}C-NMR (100 MHz, DMSO-d_6): δ = 166.9 (q, C-1), 164.3 (q, C-7), 149.9 (2C, t, C-10, C-10‘), 142.6 (q, C-5), 142.1 (q, C-8), 130.3 (2C, t, C-4, C-4‘), 126.1 (q, C-2), 121.9 (2C, t, C-9, C-9‘), 119.7 (2C, t, C-3, C-3‘) ppm; HRMS (ESI): *m/z* calc. for $C_{13}H_9N_2O_3$ $[M-H]^-$: 241.0613; found 241.0608.

4-(6-Cyanonicotinamido)-benzoic acid (43): Following the general procedure I using amine **32** (65.2 mg, 0.34 mmol) and acid **37** (100 mg, 0.68 mmol) intermediate was isolated and used directly for *t*Bu-saponification following the general procedure V**.** Acid **43** (75.0 mg, 0.28 mmol, 83%) was obtained as a beige solid.T_M = 289 °C (decomposition); ^{1}H-NMR (400 MHz, DMSO-d_6): δ = 12.7 (1H, s, CO$_2$*H*), 10.9 (1H, s, H-6), 9.23 (1H, d, *J* = 1.9 Hz, H-12), 8.53 (1H, dd, *J* = 1.9, 8.0 Hz, H-9), 8.26 (1H, d, *J* = 8.0 Hz, H-10), 7.97 (2H, m, H-4, H-4‘), 7.90 (2H, m, H-3, H-3‘) ppm; ^{13}C-NMR (100 MHz, DMSO-d_6): δ = 166.8 (q, C-1), 163.1 (q, C-7), 150.2 (t, C-12), 142.5 (q, C-5), 137.3 (t, C-9), 134.5 (q, C-8), 133.4 (q, C-11), 130.4 (2C, t, C-4, C-4‘), 128.8 (t, C-10), 126.2 (q, C-2), 119.6 (2C, t, C-3, C-3‘), 117.1 (q, CN) ppm; HRMS (ESI): *m/z* calc. for $C_{14}H_8N_3O_3$ [M – H]$^-$: 266.0566; found 266.0565.

4-(3-(4-Cyanophenyl)-propanamido)-benzoic acid (44): Following the general procedure I using amine **32** (55.1 mg, 0.29 mmol) and acid **38** (100 mg, 0.57 mmol) the intermediate was isolated and directly used for the next step, the *t*Bu-saponification following the general procedure V**.** Acid **44** (66.0 mg, 0.22 mmol, 79%) was obtained as a colourless solid. T_M = 280 °C; ^{1}H-NMR (400 MHz, DMSO-d_6): δ = 12.7 (1H, br. s, CO$_2$*H*), 10.3 (1H, s, H-6), 7.87 (2H, m, H-3, H-3‘), 7.76 (2H, m, H-12, H-12‘), 7.67 (2H, m, H-4, H-4‘), 7.47 (2H, m, H-11, H-11‘), 3.00 (2H, t, *J* = 7.5 Hz, H-9), 2.71 (2H, t, *J* = 7.5 Hz, H-8) ppm; ^{13}C-NMR (100 MHz, DMSO-d_6): δ = 170.5 (q, C-7), 166.9 (q, C-1), 147.2 (q, C-10), 143.1 (q, C-5), 132.2 (2C, t, C-12, C-12‘), 130.4 (2C, t, C-3, C-3‘), 129.4 (2C, t, C-11, C-11‘), 125.0 (q, C-2), 119,0 (q, CN), 118,3 (2C, t, C-4, C-4‘), 108.9 (q, C-13), 37.1 (s, C-8), 30.5 (s, C-9) ppm; HRMS (ESI): *m/z* calc. for $C_{17}H_{13}N_2O_3$ [M – H]$^-$: 293.0926; found 293.0924.

***tert*-Butyl-4-(2-(allyloxy)-4-(4-(4-((2*S*,3*R*)-4-amino-3-methoxy-2-(4-(4-nitro-benzamido)-benzamido)-4-oxobutanamido)phenyl)-1H-1,2,3-triazol-1-yl)-3-isopropoxybenzamido)-benzoate (S10):** Following the general procedure II acid **19** (32.0 mg, 112 µmol, 2.5 eq) and amine **31** (32.0 mg, 44.9 µmol, 1.0 eq) provided compound **S10** as a colourless amorphous solid (20.1 mg, 20.5 µmol, 45%). Preparative HPLC (RP-18; run time 100 min; H$_2$O/MeCN = 70 : 30 → 0 : 100 in 80 min; t_r = 58 min). $[\alpha]_D^{24}$ = +48.6° (*c* 2.0, DMSO); ^{1}H-NMR (400 MHz, DMSO-d_6): δ = 10.8 (1H, s, N*H*), 10.7 (1H, s, N*H*), 10.4 (1H, s, N*H*), 8.90 (1H, s, C*H*$_{\text{Triazol}}$), 8.50 (1H, d, *J* = 7.9 Hz, N*H*CH), 8.39 (2H, m, Ar*H*), 8.21 (2H, m, Ar*H*), 7.94 – 7.92 (8H, m, Ar*H*), 7.84 (2H, m, Ar*H*), 7.80 (2H, m, Ar*H*), 7.62 – 7.60 (2H, m, Ar*H*, CON*H*$_2$), 7.54 (1H, d, *J* = 8.4 Hz, Ar*H*), 7.48 (1H, s, CON*H*$_2$), 6.07 – 5.97 (1H, m, C*H*$_{\text{allyl}}$), 5.38 (1H, d, *J* = 17.2 Hz, C*H*$_{\text{allyl-}trans}$), 5.21 (1H, d, *J* = 10.3 Hz, C*H*$_{\text{allyl-}cis}$), 4.92 (1H, t, *J* = 7.9 Hz, C*H*NH), 4.69 (2H, d, *J* = 4.8 Hz, OC*H*$_2$allyl), 4.28 (1H, hept, *J* = 6.0 Hz, C*H*Me$_2$), 4.11 (1H, d, *J* = 7.9 Hz, C*H*OMe), 3.32 (3H, s, OC*H*$_3$), 1.55 (9H, s, C(C*H*$_3$)$_3$), 1.01 (6H, d, *J* = 6.0 Hz, CH(C*H*$_3$)$_2$) ppm; ^{13}C-NMR (100 MHz, DMSO-d_6): δ = 171.1 (q, CONH$_2$), 168.2 (q, *C*ONH), 165.4 (q, *C*ONH), 164.6 (q, CO$_2$*t*Bu), 164.3 (q, *C*ONH), 164.3 (q, *C*ONH), 150.3 (q, C-Ar), 149.3 (q, C-Ar), 146.3 (q, C-Ar$_{\text{Triazol}}$), 144.0 (q, C-Ar), 142.9 (q, C-Ar), 141.7 (q, C-Ar), 140.3 (q, C-Ar), 138.9 (q, C-Ar), 133.7 (q, C-Ar), 133.5 (q, C-Ar), 133.0 (t, *C*Hallyl), 130.2 (q, C-Ar), 129.4 (2C, t, C-Ar), 129.1 (q, C-Ar), 128.3 (2C, t, C-Ar), 126.3 (2C, t, C-Ar), 125.8 (2C, t, C-Ar), 125.4 (2C, t, C-Ar), 123.9 (t, C-Ar), 123.6 (2C, t, C-Ar), 122.8 (t, C-Ar$_{\text{Triazol}}$), 120.9 (t, C-Ar), 119.8 (q, C-Ar), 119.7 (2C, t, C-Ar), 118.9 (2C, t, C-Ar), 118.0 (s, *C*H$_2$allyl), 80.4 (q, *C*(CH$_3$)$_3$), 80.1 (t, *C*HOMe), 76.9 (t, *C*H(CH$_3$)$_2$), 74.6 (s, OC*H*$_2$allyl), 57.8 (p, O*C*H$_3$), 55.7 (t, *C*HNH), 27.9 (3C, p, C(*C*H$_3$)$_3$), 21.9 (2C, p, CH(*C*H$_3$)$_2$) ppm; **HRMS** (ESI): *m/z* calc. for $C_{51}H_{51}N_9O_{12}Na$ [M + Na]$^+$: 1004.3555; found 1004.3553.

***tert*-Butyl-4-(2-(allyloxy)-4-(4-(4-((2*S*,3*R*)-4-amino-2-(4-(4-cyanobenzamido)-benzamido)-3-methoxy-4-oxobutanamido)phenyl)-1H-1,2,3-triazol-1-yl)-3-isopropoxybenzamido)-benzoate (S11):** Following the general procedure II acid **20** (30.0 mg, 112 µmol, 2.5 eq) and amine **31** (32.0 mg, 44.9 µmol, 1.0 eq) provided compound **S11** as a colourless amorphous solid (19.2 mg, 20.0 µmol, 44%). Preparative HPLC (RP-18; run time 100 min; H$_2$O/MeCN = 70 : 30 → 0 : 100 in 80 min; t$_r$ = 56 min). $[\alpha]_D^{25}$ = +53.4° (*c* 1.9, DMSO); ^{1}H-NMR (400 MHz, DMSO-d$_6$): δ = 10.8 (1H, s, N*H*), 10.7 (1H, s, N*H*), 10.4 (1H, s, N*H*), 8.90 (1H, s, C*H*$_{\text{Triazol}}$), 8.51 (1H, d, *J* = 8.1 Hz, N*H*CH), 8.13 (2H, m, Ar*H*), 8.04 (2H, m, Ar*H*), 7.94 – 7.90 (8H, m, Ar*H*), 7.84 (2H, m, Ar*H*), 7.79 (2H, m, Ar*H*), 7.61 – 7.60 (2H, m, Ar*H*, CON*H*$_2$), 7.53 (1H, d, *J* = 8.3 Hz, Ar*H*), 7.47 (1H, s, CON*H*$_2$), 6.05 – 5.97 (1H, m, C*H*$_{\text{allyl}}$), 5.38 (1H, d, *J* = 17.3 Hz, C*H*$_{\text{allyl-}\mathit{trans}}$), 5.20 (1H, d, *J* = 10.5 Hz, C*H*$_{\text{allyl-}\mathit{cis}}$), 4.92 (1H, t, *J* = 8.1 Hz, C*H*NH), 4.69 (2H, d, *J* = 5.0 Hz, OC*H*$_2$allyl), 4.28 (1H, hept, *J* = 6.0 Hz, C*H*Me$_2$), 4.11 (1H, d, *J* = 8.1 Hz, C*H*OMe), 3.31 (3H, s, OC*H*$_3$), 1.55 (9H, s, C(C*H*$_3$)$_3$), 1.01 (6H, d, *J* = 6.0 Hz, CH(C*H*$_3$)$_2$) ppm; ^{13}C-NMR (100 MHz, DMSO-d$_6$): δ = 171.0 (q, CONH$_2$), 168.2 (q, *C*ONH), 165.4 (q, *C*ONH), 164.6 (q, CO$_2$*t*Bu), 164.5 (q, *C*ONH), 164.3 (q, *C*ONH), 150.3 (q, C-Ar), 146.3 (q, C-Ar$_{\text{Triazol}}$), 144.0 (q, C-Ar), 142.9 (q, C-Ar), 141.8 (q, C-Ar), 138.9 (q, C-Ar), 138.7 (q, C-Ar), 133.6 (q, C-Ar), 133.5 (q, C-Ar), 133.0 (t, *C*Hallyl), 132.5 (2C, t, C-Ar), 130.2 (q, C-Ar), 129.0 (q, C-Ar), 128.7 (2C, t, C-Ar), 128.3 (2C, t, C-Ar), 126.3 (2C, t, C-Ar), 125.8 (2C, t, C-Ar), 125.4 (2C, t, C-Ar), 123.9 (t, C-Ar), 122.8 (t, C-Ar$_{\text{Triazol}}$), 120.9 (t, C-Ar), 119.8 (q, C-Ar), 119.6 (2C, t, C-Ar), 118.9 (2C, t, C-Ar), 118.3 (q, CN), 118.0 (s, *C*H$_2$allyl), 114.1 (q, C-Ar), 80.4 (q, *C*(CH$_3$)$_3$), 80.1 (t, *C*HOMe), 76.8 (t, *C*H(CH$_3$)$_2$), 74.6 (s, OC*H*$_2$allyl), 57.7 (p, O*C*H$_3$), 55.7 (t, *C*HNH), 27.9 (3C, p, C(*C*H$_3$)$_3$), 21.9 (2C, p, CH(*C*H$_3$)$_2$) ppm; **HRMS** (ESI): *m/z* calc. for $C_{52}H_{51}N_9O_{10}Na$ [M + Na]$^+$: 984.3657; found 984.3649.

***tert*-Butyl-4-(2-(allyloxy)-4-(4-(4-((2*S*,3*R*)-4-amino-2-(4-(4-fluorobenzamido)-benzamido)-3-methoxy-4-oxobutanamido)phenyl)-1H-1,2,3-triazol-1-yl)-3-isopropoxybenzamido)-benzoate (S12):** Following the general procedure II acid **39** (43.6 mg, 168 µmol, 2.5 eq) and amine **31** (48.0 mg, 67.2 µmol, 1.0 eq) provided compound **S12** as a beige solid (33.9 mg, 35.5 µmol, 53%). Preparative HPLC (RP-18; run time 100 min; H$_2$O/MeCN = 70 : 30 → 0 : 100 in 80 min; t$_r$ = 59 min). T$_M$ = 231 °C (decomposition); $[\alpha]_D^{22}$ = +87.8° (*c* 0.5, MeOH); ^{1}H-NMR (400 MHz, DMSO-d$_6$): δ = 10.7 (1H, s, N*H*), 10.5 (1H, s, N*H*), 10.4 (1H, s, N*H*), 8.90 (1H, s, C*H*$_{\text{Triazol}}$), 8.45 (1H, d, *J* = 8.1 Hz, N*H*CH), 8.08 – 8.05

(2H, m, Ar*H*), 7.94 – 7.88 (8H, m, Ar*H*), 7.85 (2H, m, Ar*H*), 7.80 (2H, m, Ar*H*), 7.62 – 7.47 (4H, m, Ar*H*, CON*H*$_2$), 7.41 – 7.32 (2H, m, Ar*H*), 6.07 – 5.97 (1H, m, C*H*$_{\text{allyl}}$), 5.38 (1H, dd, *J* = 1.5, 17.1 Hz, C*H*$_{\text{allyl-}\textit{trans}}$), 5.21 (1H, d, *J* = 10.6 Hz, C*H*$_{\text{allyl-}\textit{cis}}$), 4.92 (1H, t, *J* = 8.1 Hz, C*H*NH), 4.69 (2H, d, *J* = 5.3 Hz, OC*H*$_2$allyl), 4.28 (1H, hept, *J* = 6.1 Hz, C*H*Me$_2$), 4.10 (1H, d, *J* = 8.1 Hz, C*H*OMe), 3.32 (3H, s, OC*H*$_3$), 1.55 (9H, s, C(C*H*$_3$)$_3$), 1.01 (6H, d, *J* = 6.1 Hz, CH(C*H*$_3$)$_2$) ppm; ^{13}C-NMR (100 MHz, DMSO-d$_6$): δ = 171.0 (q, CONH$_2$), 168.2 (q, *C*ONH), 165.4 (q, *C*ONH), 164.7 (q, CO$_2$*t*Bu), 164.6 (q, *C*ONH), 164.3 (q, *C*ONH), 164.2 (d, *J* = 254.2 Hz, *C*-F), 150.3 (q, C-Ar), 146.3 (q, C-Ar$_{\text{Triazol}}$), 143.9 (q, C-Ar), 142.8 (q, C-Ar), 142.1 (q, C-Ar), 138.9 (q, C-Ar), 133.6 (q, C-Ar), 133.5 (t, *C*Hallyl), 132.9 (q, C-Ar), 131.1 (d, *J* = 2.8 Hz, C-Ar), 130.5 (2C, d, *J* = 9.2 Hz, C-Ar), 130.1 (2C, t, C-Ar), 128.6 (q, C-Ar), 128.2 (2C, t, C-Ar), 126.3 (q, C-Ar), 125.8 (2C, t, C-Ar), 125.4 (q, C-Ar), 123.9 (t, C-Ar), 122.8 (t, C-Ar$_{\text{Triazol}}$), 120.9 (t, C-Ar), 119.8 (2C, t, C-Ar), 119.5 (2C, t, C-Ar), 118.9 (2C, t, C-Ar), 118.0 (s, *C*H$_2$allyl), 115.4 (2C, d, *J* = 21.9 Hz, C-Ar), 80.4 (q, *C*(CH$_3$)$_3$), 80.1 (t, *C*HOMe), 76.8 (t, *C*H(CH$_3$)$_2$), 74.6 (s, OC*H*$_2$allyl), 57.7 (p, O*C*H$_3$), 55.7 (t, *C*HNH), 27.8 (3C, p, C(*C*H$_3$)$_3$), 21.8 (2C, p, CH(*C*H$_3$)$_2$) ppm; HRMS (ESI): *m/z* calc. for $C_{51}H_{51}N_8O_{10}FNa$ [M + Na]$^+$: 977.3610; found 977.3611.

***tert*-Butyl-4-(2-(allyloxy)-4-(4-(4-((2*S*,3*R*)-4-amino-2-(4-(5-cyanopicolin-amido)-benzamido)-3-methoxy-4-oxobutanamido)phenyl)-1H-1,2,3-triazol-1-yl)-3-isopropoxy-benzamido)benzoate (S13):** Following the general procedure II acid **40** (31.8 mg, 119 µmol, 2.5 eq) and amine **31** (34.0 mg, 47.6 µmol, 1.0 eq) provided compound **S13** as a beige amorphous solid (23.0 mg, 23.9 µmol, 50%). Preparative HPLC (RP-18; run time 100 min; H$_2$O/MeCN = 70 : 30 → 0 : 100 in 80 min; t$_r$ = 56 min). $[\alpha]_D^{22}$ = +27.7° (*c* 0.3, MeOH); ^{1}H-NMR (400 MHz, DMSO-d$_6$): δ = 11.0 (1H, s, N*H*), 10.7 (1H, s, N*H*), 10.4 (1H, s, N*H*), 9.22 (1H, br. s, Ar*H*), 8.90 (1H, s, C*H*$_{\text{Triazol}}$), 8.60 (1H, dd, *J* = 1.7, 8.1 Hz, Ar*H*), 8.49 (1H, d, *J* = 8.1 Hz, N*H*CH), 8.31 (1H, d, *J* = 8.1 Hz, Ar*H*), 8.06 (2H, m, Ar*H*), 7.94 – 7.89 (6H, m, Ar*H*), 7.85 (2H, m, Ar*H*), 7.80 (2H, m, Ar*H*), 7.61 (1H, d, *J* = 8.3 Hz, Ar*H*), 7.54 (1H, d, *J* = 8.3 Hz, Ar*H*), 7.53 (2H, d, *J* = 40.9 Hz, CON*H*$_2$), 6.07 – 5.97 (1H, m, C*H*$_{\text{allyl}}$), 5.38 (1H, d, *J* = 17.2 Hz, C*H*$_{\text{allyl-}\textit{trans}}$), 5.20 (1H, d, *J* = 10.4 Hz, C*H*$_{\text{allyl-}\textit{cis}}$), 4.92 (1H, t, *J* = 8.1 Hz, C*H*NH), 4.69 (2H, d, *J* = 5.1 Hz, OC*H*$_2$allyl), 4.28 (1H, hept, *J* = 6.0 Hz, C*H*Me$_2$), 4.11 (1H, d, *J* = 8.1 Hz, C*H*OMe), 3.32 (3H, s, OC*H*$_3$), 1.55 (9H, s, C(C*H*$_3$)$_3$), 1.01 (6H, d, *J* = 6.0 Hz, CH(C*H*$_3$)$_2$) ppm; ^{13}C-NMR (100 MHz, DMSO-d$_6$): δ = 171.0 (q, CONH$_2$), 168.2 (q, *C*ONH), 165.4 (q, *C*ONH), 164.6 (q, CO$_2$*t*Bu), 164.3 (q, *C*ONH), 161.7 (q, *C*ONH), 152.3 (q, C-Ar), 151.5 (q, C-Ar), 150.3 (t, C-Ar), 146.3 (q, C-Ar$_{\text{Triazol}}$), 143.9 (q, C-Ar), 142.8 (q, C-Ar), 142.3 (t, C-Ar), 141.1 (q, C-Ar), 138.9 (q, C-Ar), 133.6 (q, C-Ar), 133.5 (q, C-Ar), 132.9 (t, *C*Hallyl), 130.2 (2C, t, C-Ar), 129.3 (q, C-Ar), 128.2 (2C, t, C-Ar), 126.3 (q, C-Ar), 125.8 (2C, t, C-Ar), 125.4 (q, C-Ar), 123.9 (t, C-Ar), 122.8 (t, C-Ar), 122.6 (t, C-Ar$_{\text{Triazol}}$), 120.9 (t, C-Ar), 119.9 (2C, t, C-Ar), 119.8 (2C, t, C-Ar), 118.9 (2C, t, C-Ar), 118.0 (s, *C*H$_2$allyl), 116.6 (q, CN), 111.8 (q, C-Ar), 80.4 (q, *C*(CH$_3$)$_3$), 80.0 (t, *C*HOMe), 76.8 (t, *C*H(CH$_3$)$_2$), 74.6 (s, OC*H*$_2$allyl), 57.7 (p, O*C*H$_3$), 55.7

(t, *C*HNH), 27.8 (3C, p, C(*C*H$_3$)$_3$), 21.8 (2C, p, CH(*C*H$_3$)$_2$) ppm; HRMS (ESI): *m/z* calc. for $C_{51}H_{50}N_{10}O_{10}Na$ [M + Na]$^+$: 985.3609; found 985.3609.

***tert*-Butyl-4-(2-(allyloxy)-4-(4-(4-((2*S*,3*R*)-4-amino-2-(4-(4-cyano-3-fluoro-benzamido)-benzamido)-3-methoxy-4-oxobutanamido)phenyl)-1H-1,2,3-triazol-1-yl)-3-isopropoxy-benzamido)benzoate (S14):** Following the general procedure II acid **41** (47.7 mg, 168 µmol, 2.5 eq) and amine **31** (48.0 mg, 67.2 µmol, 1.0 eq) provided compound **S14** as a colourless amorphous solid (56.2 mg, 57.4 µmol, 85%). Preparative HPLC (RP-18; run time 100 min; H$_2$O/MeCN = 70 : 30 → 0 : 100 in 80 min; t$_r$ = 59 min). $[\alpha]_D^{23}$ = +36.7° (*c* 0.3, MeOH); ^{1}H-NMR (400 MHz, DMSO-d$_6$): δ = 10.8 (1H, s, N*H*), 10.7 (1H, s, N*H*), 10.4 (1H, s, N*H*), 8.90 (1H, s, C*H*$_{\text{Triazol}}$), 8.47 (1H, d, *J* = 8.4 Hz, N*H*CH), 8.17 – 8.13 (1H, m, Ar*H*), 8.09 – 8.06 (1H, m, Ar*H*), 7.99 – 7.96 (1H, m, Ar*H*), 7.94 – 7.90 (8H, m, Ar*H*), 7.84 (2H, m, Ar*H*), 7.79 (2H, m, Ar*H*), 7.61 (1H, d, *J* = 8.3 Hz, Ar*H*), 7.56 – 7.47 (3H, m, Ar*H*, CON*H*$_2$), 6.07 – 5.97 (1H, m, C*H*$_{\text{allyl}}$), 5.38 (1H, dd, *J* = 1.5, 17.2 Hz, C*H*$_{\text{allyl-}trans}$), 5.21 (1H, dd, *J* = 1.2, 10.4 Hz, C*H*$_{\text{allyl-}cis}$), 4.92 (1H, t, *J* = 8.1 Hz, C*H*NH), 4.69 (2H, d, *J* = 5.4 Hz, OC*H*$_2$allyl), 4.28 (1H, hept, *J* = 6.1 Hz, C*H*Me$_2$), 4.09 (1H, d, *J* = 7.9 Hz, C*H*OMe), 3.31 (3H, s, OC*H*$_3$), 1.55 (9H, s, C(C*H*$_3$)$_3$), 1.01 (6H, d, *J* = 6.1 Hz, CH(C*H*$_3$)$_2$) ppm; ^{13}C-NMR (100 MHz, DMSO-d$_6$): δ = 171.0 (q, CONH$_2$), 168.1 (q, *C*ONH), 165.4 (q, *C*ONH), 164.6 (q, CO$_2$*t*Bu), 164.3 (q, *C*ONH), 163.5 (q, *C*ONH), 162.1 (d, *J* = 223.0 Hz, *C*-F), 150.3 (q, C-Ar), 146.3 (q, C-Ar$_{\text{Triazol}}$), 143.9 (q, C-Ar), 142.8 (q, C-Ar), 141.5 (d, *J* = 5.5 Hz, *C*CONH), 138.9 (q, C-Ar), 134.3 (t, C-Ar), 133.6 (q, C-Ar), 133.5 (t, *C*Hallyl), 132.9 (q, C-Ar), 130.1 (2C, t, C-Ar), 129.2 (q, C-Ar), 128.3 (2C, t, C-Ar), 126.3 (q, C-Ar), 125.8 (2C, t, C-Ar), 125.4 (q, C-Ar), 124.7 (d, *J* = 3.3 Hz, CH*C*HCCONH), 123.9 (t, C-Ar), 122.7 (t, C-Ar$_{\text{Triazol}}$), 120.9 (t, C-Ar), 120.7 (q, CN), 119.8 (2C, t, C-Ar), 119.7 (2C, t, C-Ar), 118.9 (2C, t, C-Ar), 118.0 (s, *C*H$_2$allyl), 115.8 (d, *J* = 21.8 Hz, *C*HCF), 113.6 (q, C-Ar), 102.9 (d, *J* = 15.3 Hz, *C*CN), 80.4 (q, *C*(CH$_3$)$_3$), 80.0 (t, *C*HOMe), 76.8 (t, *C*H(CH$_3$)$_2$), 74.6 (s, OC*H*$_2$allyl), 57.7 (p, O*C*H$_3$), 55.7 (t, *C*HNH), 27.8 (3C, p, C(*C*H$_3$)$_3$), 21.8 (2C, p, CH(*C*H$_3$)$_2$) ppm; HRMS (ESI): *m/z* calc. for $C_{52}H_{51}N_9O_{10}F$ [M + H]$^+$: 980.3743; found 980.3753.

***tert*-Butyl-4-(2-(allyloxy)-4-(4-(4-((2*S*,3*R*)-4-amino-2-(4-(isonicotinamido)-benzamido)-3-methoxy-4-oxobutanamido)phenyl)-1H-1,2,3-triazol-1-yl)-3-isopropoxybenzamido)-benzoate (S15):** Following to the general procedure II acid **42** (32.2 mg, 133 µmol, 2.5 eq) and amine **31** (38.0 mg, 53.2 µmol, 1.0 eq) provided compound **S15** as a colourless amorphous solid (26.7 mg, 28.5 µmol, 54%). Preparative HPLC (RP-18; run time 100 min; H$_2$O/MeCN = 70 : 30 → 0 : 100 in 80 min; t$_r$ = 50 min). $[\alpha]_D^{24}$ = +32.4° (*c* 2.3, MeOH); ^{1}H-NMR (400 MHz, DMSO-d$_6$): δ = 10.8 (1H, s, N*H*), 10.7 (1H, s, N*H*), 10.4 (1H, s, N*H*), 8.90 (1H, s, C*H*$_{\text{Triazol}}$), 8.81 (2H, d, *J* = 5.8 Hz, Ar*H*), 8.51 (1H, d, *J* = 8.1 Hz, N*H*CH), 7.94 – 7.88 (10H, m, Ar*H*), 7.85 (2H, m, Ar*H*), 7.80 (2H, m, Ar*H*), 7.62 – 7.47 (4H, m, Ar*H*, CON*H*$_2$), 6.07 – 5.97 (1H, m, C*H*$_{\text{allyl}}$), 5.38 (1H, dd, *J* = 1.6, 17.2 Hz, C*H*$_{\text{allyl-}\textit{trans}}$), 5.21 (1H, dd, *J* = 1.5, 10.5 Hz, C*H*$_{\text{allyl-}\textit{cis}}$), 4.92 (1H, t, *J* = 8.1 Hz, C*H*NH), 4.69 (2H, d, *J* = 5.4 Hz, OC*H*$_2$allyl), 4.28 (1H, hept, *J* = 6.1 Hz, C*H*Me$_2$), 4.11 (1H, d, *J* = 8.1 Hz, C*H*OMe), 3.32 (3H, s, OC*H*$_3$), 1.55 (9H, s, C(C*H*$_3$)$_3$), 1.01 (6H, d, *J* = 6.1 Hz, CH(C*H*$_3$)$_2$) ppm; ^{13}C-NMR (100 MHz, DMSO-d$_6$): δ = 171.0 (q, CONH$_2$), 168.2 (q, *C*ONH), 165.4 (q, *C*ONH), 164.6 (q, CO$_2$*t*Bu), 164.3 (q, *C*ONH), 164.3 (q, *C*ONH), 150.3 (2C, t, C-Ar), 150.3 (q, C-Ar), 146.3 (q, C-Ar$_{\text{Triazol}}$), 143.9 (q, C-Ar), 142.8 (q, C-Ar), 141.7 (q, C-Ar), 141.6 (q, C-Ar), 138.9 (q, C-Ar), 133.6 (q, C-Ar), 133.5 (t, *C*Hallyl), 132.9 (q, C-Ar), 130.1 (2C, t, C-Ar), 129.1 (q, C-Ar), 128.3 (2C, t, C-Ar), 126.3 (q, C-Ar), 125.8 (2C, t, C-Ar), 125.4 (2C, t, C-Ar), 123.9 (t, C-Ar), 122.8 (t, C-Ar$_{\text{Triazol}}$), 121.6 (2C, t, C-Ar), 120.9 (t, C-Ar), 119.8 (q, C-Ar), 119.7 (2C, t, C-Ar), 118.9 (2C, t, C-Ar), 118.0 (s, *C*H$_2$allyl), 80.4 (q, *C*(CH$_3$)$_3$), 80.1 (t, *C*HOMe), 76.8 (t, *C*H(CH$_3$)$_2$), 74.6 (s, OC*H*$_2$allyl), 57.7 (p, O*C*H$_3$), 55.7 (t, *C*HNH), 27.8 (3C, p, C(*C*H$_3$)$_3$), 21.8 (2C, p, CH(*C*H$_3$)$_2$) ppm; HRMS (ESI): *m/z* calc. for $C_{50}H_{51}N_9O_{10}Na$ [M + Na]$^+$: 960.3657; found 960.3654.

***tert*-Butyl-4-(2-(allyloxy)-4-(4-(4-((2*S*,3*R*)-4-amino-2-(4-(6-cyanonicotin-amido)-benz-amido)-3-methoxy-4-oxobutanamido)phenyl)-1H-1,2,3-triazol-1-yl)-3-isopropoxybenz-amido)benzoate (S16):** Following the general procedure II acid **43** (35.5 mg, 133 µmol, 2.5 eq) and amine **31** (38.0 mg, 53.2 µmol, 1.0 eq) provided compound **S16** as a beige amorphous solid (34.4 mg, 35.7 µmol, 67%). Preparative HPLC (RP-18; run time 100 min; H$_2$O/MeCN = 70 : 30 → 0 : 100 in 80 min; t$_r$ = 55 min). $[\alpha]_D^{23}$ = +30.5° (*c* 3.3, MeOH); ^{1}H-NMR (400 MHz, DMSO-d$_6$): δ = 10.9 (1H, s, N*H*), 10.7 (1H, s, N*H*), 10.4 (1H, s, N*H*), 9.24 (1H, d, *J* = 1.9 Hz, Ar*H*), 8.90 (1H, s, C*H*$_{\text{Triazol}}$), 8.55 (1H, dd, *J* = 1.9, 8.0 Hz, Ar*H*), 8.48 (1H, d, *J* = 8.2 Hz, N*H*CH), 8.25 (1H, d, *J* = 8.0 Hz, Ar*H*), 7.94 – 7.88 (8H, m, Ar*H*), 7.84 (2H, m, Ar*H*), 7.79 (2H, m, Ar*H*), 7.61 (1H, d, *J* = 8.3 Hz, Ar*H*), 7.54 (1H, d, *J* = 8.3 Hz, Ar*H*), 7.52 (2H, d, *J* = 39.5 Hz, CON*H*$_2$), 6.07 – 5.97 (1H, m, C*H*$_{\text{allyl}}$), 5.38 (1H, dd, *J* = 1.6, 17.2 Hz, C*H*$_{\text{allyl-}\textit{trans}}$), 5.21 (1H, dd, *J* = 1.5, 10.5 Hz, C*H*$_{\text{allyl-}\textit{cis}}$), 4.92 (1H, t, *J* = 8.1 Hz, C*H*NH), 4.69 (2H, d, *J* = 5.5 Hz, OC*H*$_2$allyl), 4.28 (1H, hept, *J* = 6.1 Hz, C*H*Me$_2$), 4.10 (1H, d, *J* = 8.0 Hz, C*H*OMe), 3.32 (3H, s, OC*H*$_3$), 1.55 (9H, s, C(C*H*$_3$)$_3$), 1.01 (6H, d, *J* = 6.1 Hz, CH(C*H*$_3$)$_2$) ppm; ^{13}C-NMR (100 MHz, DMSO-d$_6$): δ = 171.0 (q, CONH$_2$), 168.2 (q, *C*ONH), 165.4 (q, *C*ONH), 164.6 (q, CO$_2$*t*Bu), 164.3 (q, *C*ONH), 163.0 (q, *C*ONH), 150.3 (q, C-Ar), 150.2 (t, C-Ar), 146.3 (q, C-Ar$_{\text{Triazol}}$), 143.9 (q, C-Ar), 142.8 (q, C-Ar), 141.5 (q, C-Ar), 138.9 (q, C-Ar), 137.3 (t, C-Ar), 134.5 (q, C-Ar), 133.6 (q, C-Ar), 133.5 (t, *C*Hallyl), 133.4 (q, C-Ar), 132.9 (q, C-Ar), 130.1 (2C, t, C-Ar), 129.2 (q, C-Ar), 128.8 (t, C-Ar), 128.3 (2C, t, C-Ar), 126.3 (q, C-Ar), 125.8 (2C, t, C-Ar), 125.4 (q, C-Ar), 123.9 (t, C-Ar), 122.8 (t, C-Ar$_{\text{Triazol}}$), 120.9 (t, C-Ar), 119.8 (2C, t, C-Ar), 119.6 (2C, t, C-Ar), 118.9 (2C, t, C-Ar), 118.0 (s, *C*H$_2$allyl), 117.1 (q, CN), 80.4 (q, *C*(CH$_3$)$_3$), 80.0 (t, *C*HOMe), 76.8 (t, *C*H(CH$_3$)$_2$), 74.6 (s, OC*H*$_2$allyl), 57.7 (p, O*C*H$_3$), 55.7 (t, *C*HNH), 27.8 (3C, p, C(*C*H$_3$)$_3$), 21.8 (2C, p, CH(*C*H$_3$)$_2$) ppm; HRMS (ESI): *m/z* calc. for $C_{51}H_{51}N_{10}O_{10}$ [M + H]$^+$: 963.3790; found 963.3795.

***tert*-Butyl-4-(2-(allyloxy)-4-(4-(4-((2*S*,3*R*)-4-amino-2-(4-(3-(4-cyanophenyl)-propanamido)benzamido)-3-methoxy-4-oxobutanamido)phenyl)-1H-1,2,3-triazol-1-yl)-3-isopropoxybenzamido)benzoate (S17):** Following the general procedure II acid **44** (39.0 mg, 133 µmol, 2.5 eq) and amine **31** (38.0 mg, 53.2 µmol, 1.0 eq) provided phenol **S19** as a yellow amorphous solid (10.1 mg, 11.0 µmol, 56%). Preparative HPLC (RP-18; run time 100 min; H_2O/MeCN = 70 : 30 → 0 : 100 in 80 min; t_r = 57 min). $[\alpha]_D^{25}$ = +23.2° (*c* 2.7, MeOH); ^{1}H-NMR (400 MHz, DMSO-d_6): δ = 10.7 (1H, s, N*H*), 10.4 (1H, s, N*H*), 10.2 (1H, s, N*H*), 8.90 (1H, s, C*H*$_{\mathrm{Triazol}}$), 8.41 (1H, d, *J* = 8.0 Hz, N*H*CH), 7.94 – 7.91 (4H, m, Ar*H*), 7.86 – 7.75 (8H, m, Ar*H*), 7.68 (2H, m, Ar*H*), 7.61 (1H, d, *J* = 8.3 Hz, Ar*H*), 7.57 – 7.53 (2H, m, Ar*H*, CON*H*$_2$), 7.49 – 7.47 (3H, m, Ar*H*, CON*H*$_2$), 6.07 – 5.97 (1H, m, C*H*$_{\mathrm{allyl}}$), 5.38 (1H, d, *J* = 17.2 Hz, C*H*$_{\mathrm{allyl}\text{-}\mathit{trans}}$), 5.21 (1H, d, *J* = 10.4 Hz, C*H*$_{\mathrm{allyl}\text{-}\mathit{cis}}$), 4.89 (1H, t, *J* = 8.0 Hz, C*H*NH), 4.69 (2H, d, *J* = 5.1 Hz, OC*H*$_2$allyl), 4.28 (1H, hept, *J* = 6.0 Hz, C*H*Me$_2$), 4.09 (1H, d, *J* = 8.0 Hz, C*H*OMe), 3.30 (3H, s, OC*H*$_3$), 3.01 (2H, t, *J* = 7.4 Hz, ArC*H*$_2$CH$_2$), 2.71 (2H, t, *J* = 7.4 Hz, ArCH$_2$C*H*$_2$), 1.55 (9H, s, C(C*H*$_3$)$_3$), 1.01 (6H, d, *J* = 6.0 Hz, CH(C*H*$_3$)$_2$) ppm; ^{13}C-NMR (100 MHz, DMSO-d_6): δ = 171.0 (q, CONH$_2$), 170.4 (q, *C*ONH), 168.2 (q, *C*ONH), 165.4 (q, *C*ONH), 164.6 (q, CO$_2$*t*Bu), 164.3 (q, *C*ONH), 150.3 (q, C-Ar), 147.3 (q, C-Ar), 146.3 (q, C-Ar$_{\mathrm{Triazol}}$), 143.9 (q, C-Ar), 142.8 (q, C-Ar), 142.1 (q, C-Ar), 138.9 (q, C-Ar), 133.6 (q, C-Ar), 133.5 (t, *C*Hallyl), 133.0 (q, C-Ar), 132.3 (2C, t, C-Ar), 130.2 (2C, t, C-Ar), 129.5 (2C, t, C-Ar), 128.4 (2C, t, C-Ar), 128.0 (q, C-Ar), 126.3 (q, C-Ar), 125.8 (2C, t, C-Ar), 125.4 (q, C-Ar), 123.9 (t, C-Ar), 122.8 (t, C-Ar$_{\mathrm{Triazol}}$), 120.9 (t, C-Ar), 119.8 (2C, t, C-Ar), 119.0 (q, CN), 118.9 (2C, t, C-Ar), 118.2 (s, *C*H$_2$allyl), 118.0 (2C, t, C-Ar), 108.9 (q, C-Ar), 80.4 (q, *C*(CH$_3$)$_3$), 80.0 (t, *C*HOMe), 76.8 (t, *C*H(CH$_3$)$_2$), 74.6 (s, OC*H*$_2$allyl), 57.7 (p, O*C*H$_3$), 55.7 (t, *C*HNH), 37.1 (s, ArCH$_2$*C*H$_2$), 30.6 (s, Ar*C*H$_2$CH$_2$), 27.9 (3C, p, C(*C*H$_3$)$_3$), 21.9 (2C, p, CH(*C*H$_3$)$_2$) ppm; HRMS (ESI): *m/z* calc. for $C_{54}H_{56}N_9O_{10}$ $[M + H]^+$: 990.4150; found 990.4148.

***tert*-Butyl 4-(4-(4-(4-((2*S*,3*R*)-4-amino-3-methoxy-2-(4-(4-nitrobenzamido)-benzamido)-4-oxobutanamido)phenyl)-1H-1,2,3-triazol-1-yl)-2-hydroxy-3-isopropoxybenzamido)-benzoate (S18):** Following the general procedure VI compound **S10** (20.0 mg, 20.4 µmol, 1.0 eq) was used. Preparative HPLC (RP-18; run time 80 min; H_2O/MeCN = 70 : 30 → 0 : 100 in 70 min; t_r = 53 min) provided Phenol **S18** as a yellow amorphous solid (8.5 mg, 9.03 µmol, 44%). $[\alpha]_D^{24}$ = +42.9° (*c* 0.9, DMSO); ^{1}H-NMR (600 MHz, DMSO-d_6): δ = 12.2 (1H, br. s, O*H*), 11.1 (1H, br. s, N*H*), 10.8 (1H, s, N*H*), 10.4 (1H, s, N*H*), 8.92 (1H, s, C*H*$_{\mathrm{Triazol}}$), 8.44 (1H, d, *J* = 8.1 Hz, N*H*CH), 8.39 (2H, m, Ar*H*), 8.21 (2H, m, Ar*H*), 7.95 – 7.87 (11H, m, Ar*H*), 7.79 (2H, m, Ar*H*), 7.51 (2H, d, *J* = 46.5 Hz, CON*H*$_2$), 7.30 (1H, d, *J* = 8.0 Hz, Ar*H*), 4.93 (1H, t, *J* = 8.1 Hz, C*H*NH), 4.28 (1H, hept, *J* = 6.1 Hz, C*H*Me$_2$), 4.09 (1H, d, *J* = 8.1 Hz, C*H*OMe), 3.32 (3H, s, OC*H*$_3$), 1.56 (9H, s, C(C*H*$_3$)$_3$), 1.02 (6H, d, *J* = 6.1 Hz, CH(C*H*$_3$)$_2$) ppm; ^{13}C-NMR (125 MHz, DMSO-d_6): δ = 171.0 (q, CONH$_2$), 168.2 (q, *C*ONH), 167.4 (q, *C*ONH), 165.4 (q, *C*ONH), 164.5 (q, CO$_2$*t*Bu), 164.2 (q, *C*ONH), 151.5 (q, C-Ar), 149.3 (q, C-Ar), 146.2 (q, C-Ar$_{\mathrm{Triazol}}$), 142.1 (q, C-Ar), 141.7 (q, C-Ar), 140.3 (q, C-Ar), 138.9 (q, C-Ar), 138.9 (q, C-

Ar), 134.3 (q, C-Ar), 130.0 (2C, t, C-Ar), 129.3 (2C, t, C-Ar), 129.1 (q, C-Ar), 128.3 (2C, t, C-Ar), 126.8 (q, C-Ar), 125.8 (2C, t, C-Ar), 125.4 (t, C-Ar), 124.9 (q, C-Ar), 123.6 (2C, t, C-Ar), 122.6 (t, C-Ar$_{\text{Triazol}}$), 120.5 (2C, t, C-Ar), 119.8 (2C, t, C-Ar), 119.7 (2C, t, C-Ar), 118.4 (q, C-Ar), 114.1 (t, C-Ar), 80.5 (q, *C*(CH_3)$_3$), 80.1 (t, *C*HOMe), 75.4 (t, *C*H(CH_3)$_2$), 57.7 (p, O*C*H_3), 55.7 (t, *C*HNH), 27.9 (3C, p, C(*C*H_3)$_3$), 21.8 (2C, p, CH(*C*H_3)$_2$) ppm; HRMS (ESI): *m/z* calc. for $C_{48}H_{47}N_9O_{12}Na$ $[M + Na]^+$: 964.3242; found 964.3229.

***tert*-Butyl-4-(4-(4-(4-((2*S*,3*R*)-4-amino-2-(4-(4-cyanobenzamido)benzamido)-3-methoxy-4-oxobutanamido)phenyl)-1H-1,2,3-triazol-1-yl)-2-hydroxy-3-isopropoxybenzamido)-benzoate (S19):** Following the general procedure VI compound **S11** (19.0 mg, 20.4 µmol, 1.0 eq) provided phenol **S19** as a yellow amorphous solid (10.1 mg, 11.0 µmol, 56%). Preparative HPLC (RP-18; run time 80 min; H_2O/MeCN = 70 : 30 → 0 : 100 in 70 min; t_r = 52 min). $[\alpha]_D^{23}$ = +49.6° (*c* 1.0, DMSO); ^{1}H-NMR (500 MHz, DMSO-d_6): δ = 12.2 (1H, br. s, O*H*), 11.0 (1H, br. s, N*H*), 10.7 (1H, s, N*H*), 10.4 (1H, s, N*H*), 8.90 (1H, s, C*H*$_{\text{Triazol}}$), 8.44 (1H, d, *J* = 8.1 Hz, N*H*CH), 8.13 (2H, m, Ar*H*), 8.05 (2H, m, Ar*H*), 7.96 – 7.87 (11H, m, Ar*H*), 7.79 (2H, m, Ar*H*), 7.51 (2H, d, *J* = 37.9 Hz, CON*H*$_2$), 7.32 (1H, d, *J* = 8.6 Hz, Ar*H*), 4.92 (1H, t, *J* = 8.1 Hz, C*H*NH), 4.27 (1H, hept, *J* = 6.1 Hz, C*H*Me$_2$), 4.09 (1H, d, *J* = 8.1 Hz, C*H*OMe), 3.32 (3H, s, OC*H*$_3$), 1.56 (9H, s, C(C*H*$_3$)$_3$), 1.02 (6H, d, *J* = 6.1 Hz, CH(C*H*$_3$)$_2$) ppm; ^{13}C-NMR (125 MHz, DMSO-d_6): δ = 171.0 (q, CONH$_2$), 168.2 (q, *C*ONH), 167.4 (q, *C*ONH), 165.4 (q, *C*ONH), 164.5 (q, CO$_2$*t*Bu), 164.5 (q, *C*ONH), 155.0 (q, C-Ar), 146.2 (q, C-Ar$_{\text{Triazol}}$), 142.0 (q, C-Ar), 141.8 (q, C-Ar), 138.9 (q, C-Ar), 138.8 (q, C-Ar), 138.7 (q, C-Ar), 134.3 (q, C-Ar), 132.5 (2C, t, C-Ar), 130.0 (2C, t, C-Ar), 129.0 (q, C-Ar), 128.6 (2C, t, C-Ar), 128.3 (2C, t, C-Ar), 126.9 (q, C-Ar), 125.8 (2C, t, C-Ar), 125.4 (q, C-Ar), 123.6 (t, C-Ar), 122.6 (t, C-Ar$_{\text{Triazol}}$), 120.5 (2C, t, C-Ar), 119.8 (2C, t, C-Ar), 119.6 (2C, t, C-Ar), 118.4 (q, C-Ar), 118.3 (q, CN), 114.4 (q, C-Ar), 114.1 (t, C-Ar), 80.5 (q, *C*(CH_3)$_3$), 80.1 (t, *C*HOMe), 75.5 (t, *C*H(CH_3)$_2$), 57.7 (p, O*C*H_3), 55.7 (t, *C*HNH), 27.8 (3C, p, C(*C*H_3)$_3$), 21.8 (2C, p, CH(*C*H_3)$_2$) ppm; HRMS (ESI): *m/z* calc. for $C_{49}H_{48}N_9O_{10}$ $[M + H]^+$: 922.3524; found 922.3529.

***tert*-Butyl-4-(4-(4-(4-((2*S*,3*R*)-4-amino-2-(4-(4-fluorobenzamido)benzamido)-3-methoxy-4-oxobutanamido)phenyl)-1H-1,2,3-triazol-1-yl)-2-hydroxy-3-isopropoxybenzamido)-benzoate (S20):** Following the general procedure VI compound **S12** (33.9 mg, 35.5 µmol, 1.0 eq) provided Phenol **S20** as a yellow amorphous solid (20.1 mg, 22.0 µmol, 62%). Preparative HPLC (RP-18; run time 80 min; H_2O/MeCN = 70 : 30 → 0 : 100 in 70 min; t_r = 51 min). $[\alpha]_D^{24}$ = +29.2° (*c* 1.8, MeOH); ^{1}H-NMR (400 MHz, DMSO-d_6): δ = 12.2 (1H, br. s, O*H*), 11.4 (1H, s, N*H*), 10.5 (1H, s, N*H*), 10.4 (1H, s, N*H*), 8.91 (1H, s, C*H*$_{\text{Triazol}}$), 8.42 (1H, d, *J* = 8.1 Hz, N*H*CH), 8.08 – 8.05 (2H, m, Ar*H*), 7.95 – 7.86 (11H, m, Ar*H*), 7.79 (2H, m, Ar*H*), 7.51 (2H, d, *J* = 30.8 Hz, CON*H*$_2$), 7.39 (2H, m, Ar*H*), 7.25 (1H, d, *J* = 8.6 Hz, Ar*H*), 4.93 (1H, t, *J* = 8.1 Hz, C*H*NH), 4.32 (1H, hept, *J* = 6.1 Hz, C*H*Me_2), 4.10 (1H, d, *J* = 8.1 Hz, C*H*OMe), 3.32 (3H, s, OCH_3), 1.55 (9H, s, C(CH_3)$_3$), 1.01 (6H, d, *J* = 6.1 Hz, CH(CH_3)$_2$) ppm; ^{13}C-NMR (100 MHz, DMSO-d_6): δ = 171.0 (q, $CONH_2$), 168.2 (q, *C*ONH), 167.4 (q, *C*ONH), 165.0 (q, *C*ONH), 164.7 (q, CO_2*t*Bu), 164.6 (q, *C*ONH), 164.2 (d, *J* = 249.2 Hz, *C*-F), 156.0 (q, C-Ar), 146.1 (q, C-Ar$_{\text{Triazol}}$), 142.2 (q, C-Ar), 142.1 (q, C-Ar), 139.0 (q, C-Ar), 138.8 (q, C-Ar), 134.2 (q, C-Ar), 131.1 (d, *J* = 2.9 Hz, C-Ar), 130.5 (2C, d, *J* = 9.2 Hz, C-Ar), 130.0 (2C, t, C-Ar), 128.6 (q, C-Ar), 128.2 (2C, t, C-Ar), 126.7 (q, C-Ar), 125.8 (2C, t, C-Ar), 125.5 (q, C-Ar), 123.6 (t, C-Ar), 122.6 (t, C-Ar$_{\text{Triazol}}$), 120.3 (2C, t, C-Ar), 119.8 (2C, t, C-Ar), 119.5 (2C, t, C-Ar), 118.4 (q, C-Ar), 115.4 (2C, d, *J* = 21.8 Hz, C-Ar), 113.5 (t, C-Ar), 80.5 (q, *C*(CH_3)$_3$), 80.1 (t, *C*HOMe), 75.1 (t, *C*H(CH_3)$_2$), 57.7 (p, O*C*H_3), 55.6 (t, *C*HNH), 27.8 (3C, p, C(*C*H_3)$_3$), 21.8 (2C, p, CH(*C*H_3)$_2$) ppm; HRMS (ESI): *m/z* calc. for $C_{48}H_{47}N_8O_{10}FNa$ [M + Na]$^+$: 937.3297; found 937.3273.

***tert*-Butyl-4-(4-(4-(4-((2*S*,3*R*)-4-amino-2-(4-(5-cyanopicolinamido)-benz-amido)-3-methoxy-4-oxobutanamido)phenyl)-1H-1,2,3-triazol-1-yl)-2-hydroxy-3-isopropoxy-benzamido)benzoate (S21):** Following the general procedure VI compound **S13** (23.0 mg, 23.9 µmol, 1.0 eq) provided phenol **S21** as a yellow amorphous solid (21.8 mg, 23.6 µmol, 99%). Preparative HPLC (RP-18; run time 80 min; H_2O/MeCN = 70 : 30 → 0 : 100 in 70 min; t_r = 51 min). $[\alpha]_D^{24}$ = +15.4° (*c* 1.1, MeOH); ^{1}H-NMR (400 MHz, DMSO-d_6): δ = 12.2 (1H, br. s, O*H*), 11.0 (1H, s, N*H*), 10.4 (1H, s, N*H*), 9.22 (1H, br. s, Ar*H*), 8.87 (1H, s, C*H*$_{\text{Triazol}}$), 8.60 (1H, dd, *J* = 2.0, 8.1 Hz, Ar*H*), 8.44 (1H, d, *J* = 8.1 Hz, N*H*CH), 8.31 (1H, d, *J* = 8.1 Hz, Ar*H*), 8.06 (2H, m, Ar*H*), 7.92 – 7.85 (9H, m, Ar*H*), 7.78 (2H, m, Ar*H*), 7.50 (2H, d, *J* = 31.1 Hz, CONH_2), 7.08 (1H, br. s, Ar*H*), 4.92 (1H, t, *J* = 8.1 Hz, C*H*NH), 4.40 (1H, br. s, C*H*Me_2), 4.09 (1H, d, *J* = 8.1 Hz, C*H*OMe), 3.32 (3H, s, OCH_3), 1.55 (9H, s, C(CH_3)$_3$), 1.00 (6H, d, *J* = 6.1 Hz, CH(CH_3)$_2$) ppm; ^{13}C-NMR (100 MHz, DMSO-d_6): δ = 171.0 (q, $CONH_2$), 168.1 (q, *C*ONH), 167.3 (q, *C*ONH), 165.4 (q, *C*ONH), 164.6 (q, CO_2*t*Bu), 163.0 (q, C-Ar), 161.7 (q, *C*ONH), 152.3 (q, C-Ar), 151.5 (t, C-Ar), 146.0 (q, C-Ar$_{\text{Triazol}}$), 142.7 (q, C-Ar), 142.3 (t, C-Ar), 141.0 (q, C-Ar), 139.5 (q, C-Ar), 138.8 (q, C-Ar), 134.0 (q, C-Ar), 130.0 (2C, t, C-Ar), 129.3 (q, C-Ar), 128.2 (2C, t, C-Ar), 126.2 (q, C-Ar), 125.8 (2C, t, C-Ar), 125.6 (q, C-Ar), 123.7 (t, C-Ar), 122.6 (t, C-Ar), 122.6 (t, C-Ar$_{\text{Triazol}}$), 120.0 (2C, t, C-Ar), 119.9 (2C, t, C-Ar), 119.8 (2C, t, C-Ar), 118.5 (q, C-Ar), 116.6 (q, CN), 111.8 (q, C-Ar), 80.4 (q, *C*(CH_3)$_3$), 80.1 (t, *C*HOMe), 74.4 (t, *C*H(CH_3)$_2$), 57.7 (p, O*C*H_3), 55.7 (t, *C*HNH), 27.9 (3C, p, C(*C*H_3)$_3$), 21.9 (2C, p, CH(*C*H_3)$_2$) ppm; HRMS (ESI): *m/z* calc. for $C_{48}H_{46}N_{10}O_{10}Na$ [M + Na]$^+$: 945.3296; found 945.3298.

***tert*-Butyl-4-(4-(4-(4-((2*S*,3*R*)-4-amino-2-(4-(4-cyano-3-fluorobenzamido)-benzamido)-3-methoxy-4-oxobutanamido)phenyl)-1H-1,2,3-triazol-1-yl)-2-hydroxy-3-isopropoxy-**

benzamido)benzoate (S22): Following to the **general procedure VI** compound **S14** (56.2 mg, 57.4 µmol, 1.0 eq) provided phenol **S22** as a colourless amorphous solid (22.4 mg, 23.8 µmol, 42%). Preparative HPLC (RP-18; run time 80 min; H_2O/MeCN = 70 : 30 → 0 : 100 in 70 min; t_r = 50 min). $[\alpha]_D^{23}$ = +21.8° (*c* 1.1, MeOH); ^{1}H-NMR (400 MHz, DMSO-d_6): δ = 12.2 (1H, br. s, O*H*), 11.0 (1H, s, N*H*), 10.8 (1H, s, N*H*), 10.4 (1H, s, N*H*), 8.92 (1H, s, CH_{Triazol}), 8.45 (1H, d, *J* = 8.1 Hz, N*H*CH), 8.16 – 8.13 (1H, m, Ar*H*), 8.09 – 8.06 (1H, m, Ar*H*), 7.99 – 7.86 (12H, m, Ar*H*), 7.79 (2H, m, Ar*H*), 7.51 (2H, d, *J* = 30.5 Hz, CONH_2), 7.32 (1H, d, *J* = 8.6 Hz, Ar*H*), 4.93 (1H, t, *J* = 8.1 Hz, C*H*NH), 4.28 (1H, hept, *J* = 6.1 Hz, C*H*Me_2), 4.10 (1H, d, *J* = 8.1 Hz, C*H*OMe), 3.32 (3H, s, OCH_3), 1.55 (9H, s, C(CH_3)$_3$), 1.02 (6H, d, *J* = 6.1 Hz, CH(CH_3)$_2$) ppm; ^{13}C-NMR (100 MHz, DMSO-d_6): δ = 171.0 (q, CONH_2), 168.2 (q, *C*ONH), 167.4 (q, *C*ONH), 165.4 (q, *C*ONH), 164.5 (q, CO_2*t*Bu), 163.5 (q, *C*ONH), 162.1 (d, *J* = 223.6 Hz, *C*-F), 155.2 (q, C-Ar), 146.2 (q, C-Ar_{Triazol}), 142.0 (q, C-Ar), 141.5 (q, C-Ar), 141.5 (d, *J* = 7.2 Hz, *C*CONH), 141.5 (q, C-Ar), 138.9 (q, C-Ar), 138.8 (q, C-Ar), 134.3 (d, *J* = 7.0 Hz, *C*HCCN), 130.0 (2C, t, C-Ar), 129.2 (q, C-Ar), 128.3 (2C, t, C-Ar), 126.9 (q, C-Ar), 125.8 (2C, t, C-Ar), 125.4 (q, C-Ar), 124.7 (d, *J* = 3.5 Hz, CH*C*HCCONH), 123.6 (t, C-Ar), 122.6 (t, C-Ar_{Triazol}), 120.5 (2C, t, C-Ar), 119.8 (2C, t, C-Ar), 119.7 (2C, t, C-Ar), 118.3 (q, CN), 115.8 (d, *J* = 21.3 Hz, *C*HCF), 114.2 (t, C-Ar), 113.6 (q, C-Ar), 102.9 (d, *J* = 15.3 Hz, *C*CN), 80.5 (q, *C*(CH_3)$_3$), 80.1 (t, *C*HOMe), 75.5 (t, *C*H(CH_3)$_2$), 57.7 (p, O*C*H_3), 55.7 (t, *C*HNH), 27.8 (3C, p, C(*C*H_3)$_3$), 21.8 (2C, p, CH(*C*H_3)$_2$) ppm; HRMS (ESI): *m/z* calc. for $C_{49}H_{46}N_9O_{10}FNa$ $[M + Na]^+$: 962.3249; found 962.3253.

***tert*-Butyl-4-(4-(4-(4-((2*S*,3*R*)-4-amino-2-(4-(isonicotinamido)benzamido)-3-methoxy-4-oxobutanamido)phenyl)-1H-1,2,3-triazol-1-yl)-2-hydroxy-3-isopropoxybenzamido)-benzoate (S23):** Following to the general procedure VI compound **S15** (26.7 mg, 28.4 µmol, 1.0 eq) provided phenol **S23** as a yellow amorphous solid (16.5 mg, 18.4 µmol, 65%).. Preparative HPLC (RP-18; run time 80 min; H_2O/MeCN = 70 : 30 → 0 : 100 in 70 min; t_r = 45 min). $[\alpha]_D^{23}$ = +22.1° (*c* 0.9, MeOH); ^{1}H-NMR (600 MHz, DMSO-d_6): δ = 12.2 (1H, br. s, O*H*), 10.8 (1H, s, N*H*), 10.7 (1H, s, N*H*), 10.4 (1H, s, N*H*), 8.93 (1H, s, CH_{Triazol}), 8.82 (2H, m, Ar*H*), 8.45 (1H, d, *J* = 8.1 Hz, N*H*CH), 7.97 – 7.87 (13H, m, Ar*H*), 7.79 (2H, m, Ar*H*), 7.51 (2H, d, *J* = 49.8 Hz, CONH_2), 7.36 (1H, d, *J* = 8.6 Hz, Ar*H*), 4.93 (1H, t, *J* = 8.0 Hz, C*H*NH), 4.25 (1H, hept, *J* = 6.1 Hz, C*H*Me_2), 4.10 (1H, d, *J* = 8.1 Hz, C*H*OMe), 3.32 (3H, s, OCH_3), 1.56 (9H, s, C(CH_3)$_3$), 1.02 (6H, d, *J* = 6.1 Hz, CH(CH_3)$_2$) ppm; ^{13}C-NMR (125 MHz, DMSO-d_6): δ = 171.0 (q, CONH_2), 168.2 (q, *C*ONH), 167.4 (q, *C*ONH), 165.4 (q, *C*ONH), 164.5 (q, CO_2*t*Bu), 163.0 (q, *C*ONH), 154.6 (q, C-Ar), 150.4 (2C, t, C-Ar), 146.3 (q, C-Ar_{Triazol}), 141.9 (q, C-Ar), 141.6 (q, C-Ar), 138.9 (q, C-Ar), 138.7 (q, C-Ar), 134.4 (q, C-Ar), 131.5 (q, C-Ar), 130.0 (2C, t, C-Ar), 129.1 (q, C-Ar), 128.8 (2C, t, C-Ar), 127.0 (q, C-Ar), 125.8 (2C, t, C-Ar), 125.4 (q, C-Ar), 123.6 (t, C-Ar), 122.6 (t, C-Ar_{Triazol}), 121.8 (2C, t, C-Ar), 120.6 (2C, t, C-Ar), 119.8 (2C, t, C-Ar), 119.7 (2C, t, C-Ar), 118.4 (q, C-Ar), 114.7 (t, C-Ar), 80.5 (q, *C*(CH_3)$_3$), 80.1 (t, *C*HOMe), 76.7 (t, *C*H(CH_3)$_2$), 57.7

(p, O*C*H_3), 55.7 (t, *C*HNH), 27.8 (3C, p, C(*C*H_3)$_3$), 21.8 (2C, p, CH(*C*H_3)$_2$) ppm; HRMS (ESI): *m/z* calc. for $C_{47}H_{48}N_9O_{10}$ $[M + H]^+$: 898.3524; found 898.3525.

***tert*-Butyl-4-(4-(4-(4-((2*S*,3*R*)-4-amino-2-(4-(6-cyanonicotinamido)-benz-amido)-3-methoxy-4-oxobutanamido)phenyl)-1H-1,2,3-triazol-1-yl)-2-hydroxy-3-isopropoxy-benzamido)benzoate (S24):** Following to the general procedure VI compound **S16** (26.7 mg, 27.7 µmol, 1.0 eq) provided phenol **S24** as a yellow amorphous solid (24.4 mg, 26.5 µmol, 95%). Preparative HPLC (RP-18; run time 80 min; H_2O/MeCN = 70 : 30 → 0 : 100 in 70 min; t_r = 47 min). $[\alpha]_D^{23}$ = +19.5° (*c* 1.2, MeOH); ^{1}H-NMR (400 MHz, DMSO-d_6): δ = 12.2 (1H, br. s, O*H*), 11.4 (1H, br. s, N*H*), 10.9 (1H, s, N*H*), 10.4 (1H, s, N*H*), 9.24 (1H, d, *J* = 2.0 Hz, Ar*H*), 8.90 (1H, s, C*H*$_{\text{Triazol}}$), 8.55 (1H, dd, *J* = 2.0, 8.1 Hz, Ar*H*), 8.46 (1H, d, *J* = 8.2 Hz, N*H*CH), 8.25 (1H, d, *J* = 8.1 Hz, Ar*H*), 7.94 – 7.86 (11H, m, Ar*H*), 7.79 (2H, m, Ar*H*), 7.51 (2H, d, *J* = 31.3 Hz, CON*H*$_2$), 7.25 (1H, d, *J* = 8.5 Hz, Ar*H*), 4.93 (1H, t, *J* = 8.1 Hz, C*H*NH), 4.31 (1H, hept, *J* = 6.0 Hz, C*H*Me$_2$), 4.10 (1H, d, *J* = 8.1 Hz, C*H*OMe), 3.32 (3H, s, OC*H*$_3$), 1.55 (9H, s, C(C*H*$_3$)$_3$), 1.02 (6H, d, *J* = 6.0 Hz, CH(C*H*$_3$)$_2$) ppm; ^{13}C-NMR (100 MHz, DMSO-d_6): δ = 171.0 (q, CONH$_2$), 168.1 (q, *C*ONH), 167.4 (q, *C*ONH), 165.4 (q, *C*ONH), 164.6 (q, CO$_2$*t*Bu), 163.0 (q, *C*ONH), 155.9 (q, C-Ar), 150.2 (t, C-Ar), 146.1 (q, C-Ar$_{\text{Triazol}}$), 142.2 (q, C-Ar), 141.5 (q, C-Ar), 139.0 (q, C-Ar), 138.8 (q, C-Ar), 137.3 (t, C-Ar), 134.5 (q, C-Ar), 134.2 (q, C-Ar), 133.4 (q, C-Ar), 130.0 (2C, t, C-Ar), 129.2 (q, C-Ar), 128.8 (t, C-Ar), 128.3 (2C, t, C-Ar), 126.7 (q, C-Ar), 125.8 (2C, t, C-Ar), 125.5 (q, C-Ar), 123.6 (t, C-Ar), 122.6 (t, C-Ar$_{\text{Triazol}}$), 120.4 (2C, t, C-Ar), 119.8 (2C, t, C-Ar), 119.6 (2C, t, C-Ar), 118.4 (q, C-Ar), 117.1 (q, CN), 113.5 (t, C-Ar), 80.5 (q, *C*(CH$_3$)$_3$), 80.1 (t, *C*HOMe), 75.2 (t, *C*H(CH$_3$)$_2$), 57.7 (p, O*C*H$_3$), 55.7 (t, *C*HNH), 27.9 (3C, p, C(*C*H$_3$)$_3$), 21.8 (2C, p, CH(*C*H$_3$)$_2$) ppm; HRMS (ESI): *m/z* calc. for $C_{48}H_{47}N_{10}O_{10}$ $[M + H]^+$: 923.3477; found 923.3469.

***tert*-Butyl-4-(4-(4-(4-((2*S*,3*R*)-4-amino-2-(4-(3-(4-cyanophenyl)-propan-amido)-benz-amido)-3-methoxy-4-oxobutanamido)phenyl)-1H-1,2,3-triazol-1-yl)-2-hydroxy-3-isopropoxybenzamido)benzoate (S25):** Following to the general procedure VI compound **S17** (29.5 mg, 29.8 µmol, 1.0 eq) provided Phenol **S25** as a yellow amorphous solid (19.8 mg, 20.9 µmol, 70%). Preparative HPLC (RP-18; run time 80 min; H_2O/MeCN = 70 : 30 → 0 : 100 in 70 min; t_r = 49 min). $[\alpha]_D^{22}$ = +14.9° (*c* 1.0, MeOH); ^{1}H-NMR (400 MHz, DMSO-d_6): δ = 12.2 (1H, br. s, O*H*), 11.2 (1H, s, N*H*), 10.4 (1H, s, N*H*), 10.2 (1H, s, N*H*), 8.92 (1H, s, C*H*$_{\text{Triazol}}$), 8.37 (1H, d, *J* = 8.0 Hz, N*H*CH), 7.95 – 7.87 (7H, m, Ar*H*), 7.83 – 7.75 (6H, m, Ar*H*), 7.68 (2H, m, Ar*H*), 7.54 – 7.46 (4H, m, Ar*H*, CON*H*$_2$), 7.29 (1H, d, *J* = 8.5 Hz, Ar*H*), 4.90 (1H, t, *J* = 8.0 Hz, C*H*NH), 4.29 (1H, hept, *J* = 6.1 Hz, C*H*Me$_2$), 4.08 (1H, d, *J* = 8.0 Hz, C*H*OMe), 3.31 (3H, s, OC*H*$_3$), 3.01 (2H, t, *J* = 7.4 Hz, ArC*H*$_2$CH$_2$), 2.71 (2H, t, *J* = 7.5 Hz, ArCH$_2$C*H*$_2$), 1.55 (9H, s, C(C*H*$_3$)$_3$), 1.02 (6H, d, *J* = 6.1 Hz, CH(C*H*$_3$)$_2$) ppm; ^{13}C-NMR (100 MHz, DMSO-d_6): δ = 171.1 (q, CONH$_2$), 170.5 (q, *C*ONH), 168.2 (q, *C*ONH), 167.4 (q, *C*ONH), 165.5 (q, *C*ONH), 164.6 (q, CO$_2$*t*Bu), 147.3 (q, C-Ar), 146.2 (q, C-Ar$_{\text{Triazol}}$), 142.1 (q, C-Ar), 139.0 (q, C-Ar), 138.9 (q, C-Ar), 134.3 (q, C-Ar), 132.3 (2C, t, C-Ar), 130.0 (2C, t, C-Ar), 129.5 (2C, t, C-Ar), 128.4 (2C, t, C-Ar), 128.1 (q, C-Ar), 126.9 (q, C-Ar), 125.9 (2C, t, C-Ar), 125.5 (q, C-Ar), 123.6 (t, C-Ar), 122.6 (t, C-Ar$_{\text{Triazol}}$), 120.5 (2C, t, C-Ar), 119.9 (2C, t, C-Ar), 119.0 (q, C-Ar), 118.5 (q, CN), 118.3 (2C, t, C-Ar), 108.9 (q, C-Ar), 80.6 (q, *C*(CH$_3$)$_3$), 80.1 (t, *C*HOMe), 75.4 (t, *C*H(CH$_3$)$_2$), 57.8 (p, O*C*H$_3$), 55.7 (t, *C*HNH), 37.2 (s, ArCH$_2$*C*H$_2$), 30.6 (s, Ar*C*H$_2$CH$_2$), 27.9 (3C, p, C(*C*H$_3$)$_3$), 21.8 (2C, p, CH(*C*H$_3$)$_2$) ppm; HRMS (ESI): *m/z* calc. for $C_{51}H_{51}N_9O_{10}Na$ [M + Na]$^+$: 972.3657; found 972.3661.

4-(4-(4-(4-((2*S*,3*R*)-4-Amino-3-methoxy-2-(4-(4-nitrobenzamido)benzamido)-4-oxo-butanamido)phenyl)-1H-1,2,3-triazol-1-yl)-2-hydroxy-3-isopropoxy-benzamido)-benzoic acid (45): Following to the general procedure V Phenol **S18** (8.5 mg, 9.03 µmol, 1.0 eq) was stirred for 6 h. Acid **45** was obtained as a colourless amorphous solid (6.2 mg, 7.00 µmol, 78%). $[\alpha]_D^{24}$ = +37.1° (*c* 0.1, DMSO); ^{1}H-NMR (500 MHz, DMSO-d$_6$): δ = 12.9 (1H, br. s, CO$_2$*H*), 12.2 (1H, br. s, O*H*), 10.8 (1H, br. s, N*H*), 10.8 (1H, s, N*H*), 10.4 (1H, s, N*H*), 8.93 (1H, s, C*H*$_{\text{Triazol}}$), 8.44 (1H, d, *J* = 8.1 Hz, N*H*CH), 8.39 (2H, m, Ar*H*), 8.21 (2H, m, Ar*H*), 8.00 – 7.87 (11H, m, Ar*H*), 7.79 (2H, m, Ar*H*), 7.50 (2H, d, *J* = 39.2 Hz, CON*H*$_2$), 7.37 (1H, d, *J* = 8.6 Hz, Ar*H*), 4.92 (1H, t, *J* = 8.1 Hz, C*H*NH), 4.24 (1H, hept, *J* = 6.1 Hz, C*H*Me$_2$), 4.09 (1H, d, *J* = 8.1 Hz, C*H*OMe), 3.32 (3H, s, OC*H*$_3$), 1.02 (6H, d, *J* = 6.1 Hz, CH(C*H*$_3$)$_2$) ppm; ^{13}C-NMR (125 MHz, DMSO-d$_6$): δ = 171.0 (q, CONH$_2$), 168.2 (q, *C*ONH), 167.5 (q, *C*ONH), 166.9 (q, *C*O$_2$H), 165.4 (q, *C*ONH), 164.2 (q, *C*ONH), 154.5 (q, C-Ar), 149.3 (q, C-Ar), 146.3 (q, C-Ar$_{\text{Triazol}}$), 141.9 (q, C-Ar), 141.7 (q, C-Ar), 140.3 (q, C-Ar), 138.9 (q, C-Ar), 138.7 (q, C-Ar), 134.4 (q, C-Ar), 130.7 (2C, t, C-Ar), 130.3 (q, C-Ar), 129.4 (2C, t, C-Ar), 128.7 (2C, t, C-Ar), 128.3 (q, C-Ar), 126.5 (q, C-Ar), 125.8 (2C, t, C-Ar), 125.4 (t, C-Ar), 123.6 (2C, t, C-Ar), 122.6 (t, C-Ar$_{\text{Triazol}}$), 120.6 (2C, t, C-Ar), 119.8 (2C, t, C-Ar), 119.7 (2C, t, C-Ar), 118.4 (q, C-Ar), 114.8 (t, C-Ar), 80.1 (t, *C*HOMe), 75.7 (t, *C*H(CH$_3$)$_2$), 57.7 (p, O*C*H$_3$), 55.7 (t, *C*HNH), 21.8 (2C, p, CH(*C*H$_3$)$_2$) ppm; HRMS (ESI): *m/z* calc. for $C_{44}H_{40}N_9O_{12}$ [M + H]$^+$: 886.2796; found 886.2798.

4-(4-(4-(4-((2*S*,3*R*)-4-Amino-2-(4-(4-cyanobenzamido)benzamido)-3-methoxy-4-oxo-butanamido)phenyl)-1H-1,2,3-triazol-1-yl)-2-hydroxy-3-iso-propoxybenzamido)-benzoic acid (46): Following the general procedure V Phenol **S19** (10.1 mg, 11.0 µmol, 1.0 eq) was stirred for 3 h. Acid **46** was obtained as a beige amorphous solid (7.4 mg, 8.55 µmol, 78%). $[\alpha]_D^{25}$ = +55.9° (*c* 0.2, DMSO); ^{1}H-NMR (600 MHz, DMSO-d$_6$): δ = 12.8 (1H, br. s, CO$_2$*H*), 12.2 (1H, br. s, O*H*), 10.8 (1H, br. s, N*H*), 10.7 (1H, s, N*H*), 10.4 (1H, s, N*H*), 8.93 (1H, s, C*H*$_{\text{Triazol}}$), 8.43 (1H, d, *J* = 8.1 Hz, N*H*CH), 8.13 (2H, m, Ar*H*), 8.05 (2H, m, Ar*H*), 7.99 – 7.87 (11H, m, Ar*H*), 7.79 (2H, m, Ar*H*), 7.50 (2H, d, *J* = 45.0 Hz, CON*H*$_2$), 7.34 (1H, d, *J* = 7.0 Hz, Ar*H*), 4.92 (1H, t, *J* = 8.1 Hz, C*H*NH), 4.26 (1H, hept, *J* = 6.1 Hz, C*H*Me$_2$), 4.09 (1H, d, *J* = 8.1 Hz, C*H*OMe), 3.32 (3H, s, OC*H*$_3$), 1.02 (6H, d, *J* = 6.1 Hz, CH(C*H*$_3$)$_2$) ppm; ^{13}C-NMR (125 MHz, DMSO-d$_6$): δ = 171.0 (q, CONH$_2$), 168.2 (q, *C*ONH), 167.4 (q, *C*ONH), 166.8 (q, *C*O$_2$H), 165.4 (q, *C*ONH), 164.5 (q, *C*ONH), 154.7 (q, C-Ar), 146.2 (q, C-Ar$_{\mathrm{Triazol}}$), 141.9 (q, C-Ar), 141.7 (q, C-Ar), 138.9 (q, C-Ar), 138.8 (q, C-Ar), 138.7 (q, C-Ar), 134.3 (q, C-Ar), 132.5 (2C, t, C-Ar), 130.3 (2C, t, C-Ar), 129.0 (q, C-Ar), 128.6 (2C, t, C-Ar), 128.3 (2C, t, C-Ar), 126.4 (q, C-Ar), 125.8 (2C, t, C-Ar), 125.4 (q, C-Ar), 123.5 (t, C-Ar), 122.6 (t, C-Ar$_{\mathrm{Triazol}}$), 120.6 (2C, t, C-Ar), 119.8 (2C, t, C-Ar), 119.6 (2C, t, C-Ar), 118.4 (q, C-Ar), 118.3 (q, CN), 114.6 (q, C-Ar), 114.0 (t, C-Ar), 80.1 (t, *C*HOMe), 75.6 (t, *C*H(CH$_3$)$_2$), 57.7 (p, O*C*H$_3$), 55.6 (t, *C*HNH), 21.8 (2C, p, CH(*C*H$_3$)$_2$) ppm; HRMS (ESI): *m/z* calc. for $C_{45}H_{40}N_9O_{10}$ [M + H]$^+$: 866.2898; found 866.2897.

4-(4-(4-(4-((2*S*,3*R*)-4-Amino-2-(4-(4-fluorobenzamido)benzamido)-3-methoxy-4-oxo-butanamido)phenyl)-1H-1,2,3-triazol-1-yl)-2-hydroxy-3-iso-propoxybenzamido)-benzoic acid (47): Following the general procedure V phenol **S20** (20.1 mg, 22.0 µmol, 1.0 eq) was stirred for 3.5 h. Acid **47** was obtained as an orange amorphous solid (19.1 mg, 19.6 µmol, 89%). **[*α*]$_D^{24}$** = +27.6° (*c* 0.9, DMSO); **^{1}H-NMR** (600 MHz, DMSO-d$_6$): δ = 12.2 (1H, br. s, O*H*), 10.8 (1H, s, N*H*), 10.5 (1H, s, N*H*), 10.4 (1H, s, N*H*), 8.93 (1H, s, C*H*$_{\mathrm{Triazol}}$), 8.41 (1H, d, *J* = 8.1 Hz, N*H*CH), 8.07 – 8.05 (2H, m, Ar*H*), 8.00 – 7.87 (11H, m, Ar*H*), 7.79 (2H, m, Ar*H*), 7.50 (2H, d, *J* = 47.2 Hz, CON*H*$_2$), 7.40 – 7.36 (3H, m, Ar*H*), 4.92 (1H, t, *J* = 8.1 Hz, C*H*NH), 4.25 (1H, hept, *J* = 6.1 Hz, C*H*Me$_2$), 4.09 (1H, d, *J* = 8.1 Hz, C*H*OMe), 3.32 (3H, s, OC*H*$_3$), 1.03 (6H, d, *J* = 6.1 Hz, CH(C*H*$_3$)$_2$) ppm; **^{13}C-NMR** (125 MHz, DMSO-d$_6$): δ = 171.0 (q, CONH$_2$), 168.2 (q, *C*ONH), 167.5 (q, *C*ONH), 166.9 (q, *C*O$_2$H), 165.0 (q, *C*ONH), 164.7 (q, *C*ONH), 164.4 (d, *J* = 311.0 Hz, *C*-F), 158.2 (TFA), 154.5 (q, C-Ar), 146.3 (q, C-Ar$_{\mathrm{Triazol}}$), 142.1 (q, C-Ar), 141.9 (q, C-Ar), 138.9 (q, C-Ar), 138.7 (q, C-Ar), 134.4 (q, C-Ar), 131.1 (d, *J* = 2.8 Hz, C-Ar), 130.5 (2C, d, *J* = 9.1 Hz, C-Ar), 130.3 (2C, t, C-Ar), 129.8 (TFA), 128.6 (q, C-Ar), 128.2 (2C, t, C-Ar), 126.5 (q, C-Ar), 125.8 (2C, t, C-Ar), 125.4 (q, C-Ar), 123.5 (t, C-Ar), 122.6 (t, C-Ar$_{\mathrm{Triazol}}$), 120.6 (2C, t, C-Ar), 119.8 (2C, t, C-Ar), 119.5 (2C, t, C-Ar), 118.4 (q, C-Ar), 115.4 (2C, d, *J* = 21.8 Hz, C-Ar), 114.8 (t, C-Ar), 80.1 (t, *C*HOMe), 75.7 (t, *C*H(CH$_3$)$_2$), 57.7 (p, O*C*H$_3$), 55.6 (t, *C*HNH), 21.8 (2C, p, CH(*C*H$_3$)$_2$) ppm; HRMS (ESI): *m/z* calc. for $C_{44}H_{40}N_8O_{10}F$ [M + H]$^+$: 859.2851; found 859.2851.

4-(4-(4-(4-((2*S*,3*R*)-4-Amino-2-(4-(5-cyanopicolinamido)benzamido)-3-methoxy-4-oxo-butanamido)phenyl)-1H-1,2,3-triazol-1-yl)-2-hydroxy-3-iso-propoxybenzamido)benzoic acid (48): Following the general procedure V phenol **S21** (21.8 mg, 23.6 µmol, 1.0 eq) was stirred for 6 h. Acid **48** was obtained as a yellow amorphous solid (8.2 mg, 9.47 µmol, 40%). [*α*]$_D^{24}$ = +27.2° (*c* 0.8, DMSO); ^{1}H-NMR (600 MHz, DMSO-d$_6$): δ = 12.9 (1H, br. s, CO$_2$*H*), 12.2 (1H, br. s, O*H*), 11.0 (1H, s, N*H*), 10.8 (1H, s, N*H*), 10.4 (1H, s, N*H*), 9.22 (1H, dd, *J* = 0.7, 2.0 Hz, Ar*H*), 8.93 (1H, s, C*H*$_{\mathrm{Triazol}}$), 8.61 (1H, dd, *J* = 2.0, 8.2 Hz, Ar*H*), 8.43 (1H, d, *J* = 8.1 Hz, N*H*CH), 8.31 (1H, dd, *J* = 0.7, 8.2 Hz, Ar*H*), 8.06 (2H, m, Ar*H*), 7.99 – 7.87 (9H, m, Ar*H*), 7.79 (2H, m, Ar*H*), 7.50 (2H, d, *J* = 43.0 Hz, CONH_2), 7.37 (1H, d, *J* = 8.6 Hz, Ar*H*), 4.91 (1H, t, *J* = 8.1 Hz, C*H*NH), 4.24 (1H, hept, *J* = 6.1 Hz, C*H*Me$_2$), 4.09 (1H, d, *J* = 8.1 Hz, C*H*OMe), 3.32 (3H, s, OCH_3), 1.02 (6H, d, *J* = 6.1 Hz, CH(CH_3)$_2$) ppm; ^{13}C-NMR (125 MHz, DMSO-d$_6$): δ = 171.0 (q, CONH$_2$), 168.2 (q, *C*ONH), 167.5 (q, *C*ONH), 166.8 (q, *C*O$_2$H), 165.4 (q, *C*ONH), 161.7 (q, *C*ONH), 154.5 (q, C-Ar), 152.3 (q, C-Ar), 151.5 (t, C-Ar), 146.3 (q, C-Ar$_{\mathrm{Triazol}}$), 142.3 (t, C-Ar), 141.9 (q, C-Ar), 141.0 (q, C-Ar), 138.9 (q, C-Ar), 138.7 (q, C-Ar), 134.4 (q, C-Ar), 130.3 (2C, t, C-Ar), 129.3 (q, C-Ar), 128.2 (2C, t, C-Ar), 126.5 (q, C-Ar), 125.8 (2C, t, C-Ar), 125.4 (q, C-Ar), 123.5 (t, C-Ar), 122.6 (t, C-Ar), 122.6 (t, C-Ar$_{\mathrm{Triazol}}$), 120.6 (2C, t, C-Ar), 119.9 (2C, t, C-Ar), 119.8 (2C, t, C-Ar), 118.4 (q, C-Ar), 116.7 (q, CN), 114.8 (t, C-Ar), 111.8 (q, C-Ar), 80.1 (t, *C*HOMe), 75.7 (t, *C*H(CH$_3$)$_2$), 57.7 (p, O*C*H$_3$), 55.6 (t, *C*HNH), 21.8 (2C, p, CH(*C*H$_3$)$_2$) ppm; HRMS (ESI): *m/z* calc. for $C_{44}H_{39}N_{10}O_{10}$ [M + H]$^+$: 867.2851; found 867.2857.

4-(4-(4-(4-((2*S*,3*R*)-4-Amino-2-(4-(4-cyano-3-fluorobenzamido)benzamido)-3-methoxy-4-oxobutanamido)phenyl)-1H-1,2,3-triazol-1-yl)-2-hydroxy-3-isopropoxybenzamido)-benzoic acid (49): Following the general procedure V phenol **S22** (22.4 mg, 23.8 µmol, 1.0 eq) was stirred for 2 h. Acid **49** was obtained as a beige amorphous solid (21.0 mg, 23.8 µmol, quant.). $[\alpha]_D^{24}$ = +39.5° (*c* 1.3, DMSO); ^{1}H-NMR (600 MHz, DMSO-d$_6$): δ = 12.8 (1H, br. s, CO$_2$*H*), 12.2 (1H, br. s, O*H*), 10.8 (1H, s, N*H*), 10.8 (1H, s, N*H*), 10.4 (1H, s, N*H*), 8.93 (1H, s, CH_{Triazol}), 8.45 (1H, d, *J* = 8.1 Hz, N*H*CH), 8.16 – 8.13 (1H, m, Ar*H*), 8.09 – 8.07 (1H, m, Ar*H*), 8.00 – 7.96 (4H, m, Ar*H*), 7.93 – 7.87 (8H, m, Ar*H*), 7.79 (2H, m, Ar*H*), 7.51 (2H, d, *J* = 47.9 Hz, CONH_2), 7.37 (1H, d, *J* = 8.6 Hz, Ar*H*), 4.92 (1H, t, *J* = 8.1 Hz, C*H*NH), 4.25 (1H, hept, *J* = 6.1 Hz, C*H*Me$_2$), 4.09 (1H, d, *J* = 8.1 Hz, C*H*OMe), 3.32 (3H, s, OCH_3), 1.03 (6H, d, *J* = 6.1 Hz, CH(CH_3)$_2$) ppm; ^{13}C-NMR (125 MHz, DMSO-d$_6$): δ = 171.0 (q, CONH$_2$), 168.2 (q, *C*ONH), 167.5 (q, *C*ONH), 166.9 (q, *C*O$_2$H), 165.4 (q, *C*ONH), 163.2 (q, *C*ONH), 162.2 (d, *J* = 256.3 Hz, *C*-F), 154.5 (q, C-Ar), 146.3 (q, C-Ar$_{\mathrm{Triazol}}$), 141.9 (q, C-Ar), 141.5 (d, *J* = 6.1 Hz, *C*CONH), 138.9 (q, C-Ar), 138.7 (q, C-Ar), 134.4 (d, *J* = 7.0 Hz, *C*HCCN), 130.7 (q, C-Ar), 130.3 (2C, t, C-Ar), 129.2 (q, C-Ar), 128.8 (q, C-Ar), 128.3 (2C, t, C-Ar), 126.5 (q, C-Ar), 125.8 (2C, t, C-Ar), 125.4 (q, C-Ar), 124.7 (d, *J* = 3.4 Hz, CH*C*HCCONH), 123.5 (t, C-Ar), 122.6 (t, C-Ar$_{\mathrm{Triazol}}$), 120.6 (2C, t, C-Ar), 119.8 (2C, t, C-Ar), 119.7 (2C, t, C-Ar), 118.4 (q, CN), 115.8 (d, *J* = 21.3 Hz, *C*HCF), 114.8 (t, C-Ar), 113.6 (q, C-Ar), 102.9 (d, *J* = 15.3 Hz, *C*CN), 80.1 (t, *C*HOMe), 75.7 (t, *C*H(CH$_3$)$_2$), 57.7 (p, O*C*H$_3$), 55.7 (t, *C*HNH), 21.8 (2C, p, CH(*C*H$_3$)$_2$) ppm; HRMS (ESI): *m/z* calc. for $C_{45}H_{39}N_9O_{10}F$ [M + H]$^+$: 884.2804; found 884.2808.

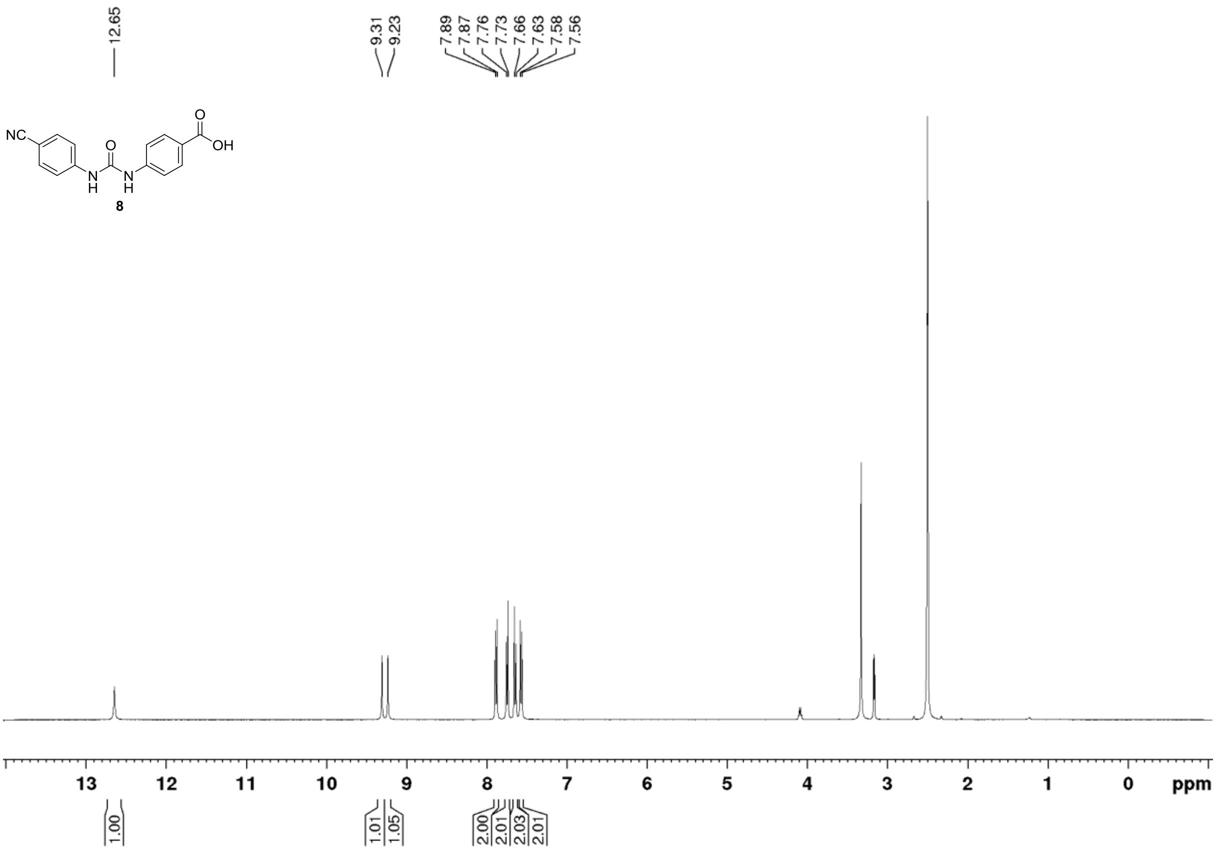
4-(4-(4-(4-((2*S*,3*R*)-4-Amino-2-(4-(isonicotinamido)benzamido)-3-methoxy-4-oxo-butanamido)phenyl)-1H-1,2,3-triazol-1-yl)-2-hydroxy-3-isopropoxy-benzamido)-benzoic acid (50): Following the general procedure V phenol **S23** (16.5 mg, 18.4 µmol, 1.0 eq) was stirred for 4 h. Acid **50** was obtained as a yellow amorphous solid (13.7 mg, 16.3 µmol, 89%). $[\alpha]_D^{24}$ = +22.6° (*c* 1.4, DMSO); ^{1}H-NMR (500 MHz, DMSO-d$_6$): δ = 12.2 (1H, br. s, O*H*), 10.8 (1H, s, N*H*), 10.8 (1H, s, N*H*), 10.4 (1H, s, N*H*), 8.93 (1H, s, C*H*$_{\text{Triazol}}$), 8.85 (2H, m, Ar*H*), 8.44 (1H, d, *J* = 8.0 Hz, N*H*CH), 8.13 – 7.87 (13H, m, Ar*H*), 7.79 (2H, m, Ar*H*), 7.50 (2H, d, *J* = 31.9 Hz, CON*H*$_2$), 7.37 (1H, d, *J* = 8.6 Hz, Ar*H*), 4.92 (1H, t, *J* = 8.0 Hz, C*H*NH), 4.25 (1H, hept, *J* = 6.1 Hz, C*H*Me$_2$), 4.09 (1H, d, *J* = 8.0 Hz, C*H*OMe), 3.32 (3H, s, OC*H*$_3$), 1.03 (6H, d, *J* = 6.1 Hz, CH(C*H*$_3$)$_2$) ppm; ^{13}C-NMR (125 MHz, DMSO-d$_6$): δ = 171.0 (q, CONH$_2$), 168.2 (q, *C*ONH), 167.5 (q, *C*ONH), 166.9 (q, *C*O$_2$H), 165.4 (q, *C*ONH), 164.1 (q, *C*ONH), 154.5 (q, C-Ar), 149.8 (2C, t, C-Ar), 146.3 (q, C-Ar$_{\text{Triazol}}$), 141.9 (q, C-Ar), 141.5 (q, C-Ar), 138.9 (q, C-Ar), 138.7 (q, C-Ar), 134.4 (q, C-Ar), 130.3 (2C, t, C-Ar), 129.2 (q, C-Ar), 128.3 (2C, t, C-Ar), 126.5 (q, C-Ar), 125.9 (q, C-Ar),125.8 (2C, t, C-Ar), 125.4 (q, C-Ar), 123.5 (t, C-Ar), 122.6 (t, C-Ar$_{\text{Triazol}}$), 122.0 (2C, t, C-Ar), 120.6 (2C, t, C-Ar), 119.8 (2C, t, C-Ar), 119.7 (2C, t, C-Ar), 118.4 (q, C-Ar), 114.8 (t, C-Ar), 80.1 (t, *C*HOMe), 75.7 (t, *C*H(CH$_3$)$_2$), 57.7 (p, O*C*H$_3$), 55.7 (t, *C*HNH), 21.8 (2C, p, CH(*C*H$_3$)$_2$) ppm; HRMS (ESI): *m/z* calc. for $C_{43}H_{40}N_9O_{10}$ [M + H]$^+$: 842.2898; found 842.2903.

4-(4-(4-(4-((2*S*,3*R*)-4-Amino-2-(4-(6-cyanonicotinamido)benzamido)-3-methoxy-4-oxo-butanamido)phenyl)-1H-1,2,3-triazol-1-yl)-2-hydroxy-3-iso-propoxybenzamido)benzoic acid (51): Following the general procedure V phenol **S24** (24.4 mg, 26.5 µmol, 1.0 eq) was stirred for 5 h. Acid **51** was obtained as a beige amorphous solid (15.6 mg, 18.0 µmol, 68%). $[\alpha]_D^{22}$ = +25.1° (*c* 1.5, DMSO); ^{1}H-NMR (600 MHz, DMSO-d$_6$): δ = 12.8 (1H, br. s, CO$_2$*H*), 12.2 (1H, br. s, O*H*), 10.9 (1H, br. s, N*H*), 10.8 (1H, s, N*H*), 10.4 (1H, s, N*H*), 9.24 – 9.23 (1H, m, Ar*H*), 8.93 (1H, s, C*H*$_{\text{Triazol}}$), 8.55 – 8.53 (1H, m, Ar*H*), 8.45 (1H, d, *J* = 8.1 Hz, N*H*CH), 8.26 – 8.25 (1H, m, Ar*H*), 8.00 – 7.87 (11H, m, Ar*H*), 7.79 (2H, m, Ar*H*), 7.50 (2H, d, *J* = 46.3 Hz, CON*H*$_2$), 7.37 (1H, d, *J* = 8.6 Hz, Ar*H*), 4.92 (1H, t, *J* = 8.1 Hz, C*H*NH), 4.25 (1H, hept, *J* = 6.0 Hz, C*H*Me$_2$), 4.09 (1H, d, *J* = 8.1 Hz, C*H*OMe), 3.32 (3H, s, OC*H*$_3$), 1.03 (6H, d, *J* = 6.0 Hz, CH(C*H*$_3$)$_2$) ppm; ^{13}C-NMR (125 MHz, DMSO-d$_6$): δ = 171.0 (q, CONH$_2$), 168.2 (q, *C*ONH), 167.5 (q, *C*ONH), 166.8 (q, *C*O$_2$H), 165.3 (q, *C*ONH), 163.0 (q, *C*ONH), 154.5 (q, C-Ar), 150.2 (t, C-Ar), 146.3 (q, C-Ar$_{\text{Triazol}}$), 141.9 (q, C-Ar), 141.5 (q, C-Ar), 138.9 (q, C-Ar), 138.7 (q, C-Ar), 137.3 (t, C-Ar), 134.5 (q, C-Ar), 134.4 (q, C-Ar), 133.4 (q, C-Ar), 130.3 (2C, t, C-Ar), 129.2 (q, C-Ar), 128.8 (t, C-Ar), 128.3 (2C, t, C-Ar), 126.5 (q, C-Ar), 125.8 (2C, t, C-Ar), 125.4 (q, C-Ar), 123.5 (t, C-Ar), 122.6 (t, C-Ar$_{\text{Triazol}}$), 120.6 (2C, t, C-Ar), 119.8 (2C, t, C-Ar), 119.6 (2C, t, C-Ar), 118.4 (q, C-Ar), 117.1 (q, CN), 114.8 (t, C-Ar), 80.0 (t, *C*HOMe), 75.7 (t, *C*H(CH$_3$)$_2$), 57.7 (p, O*C*H$_3$), 55.6 (t, *C*HNH), 21.8 (2C, p, CH(*C*H$_3$)$_2$) ppm; HRMS (ESI): *m/z* calc. for $C_{44}H_{39}N_{10}O_{10}$ [M + H]$^+$: 867.2851; found 867.2858.

4-(4-(4-(4-((2*S*,3*R*)-4-Amino-2-(4-(3-(4-cyanophenyl)-propanamido)-benz-amido)-3-methoxy-4-oxobutanamido)phenyl)-1H-1,2,3-triazol-1-yl)-2-hydroxy-3-isopropoxy-benzamido)benzoic acid (52): Following the general procedure V phenol **S25** (19.8 mg, 20.9 µmol, 1.0 eq) was stirred for 4.5 h.

Acid **52** was obtained as a grey amorphous solid (18.6 mg, 20.9 µmol, quant.). $[\alpha]_D^{24}$ = +20.8° (*c* 1.9, DMSO); ^{1}H-NMR (400 MHz, DMSO-d_6): δ = 12.8 (1H, br. s, $\mathrm{CO_2}H$), 12.2 (1H, br. s, O*H*), 10.8 (1H, s, N*H*), 10.3 (1H, s, N*H*), 10.2 (1H, s, N*H*), 8.92 (1H, s, $\mathrm{C}H_{\mathrm{Triazol}}$), 8.35 (1H, d, *J* = 8.1 Hz, N*H*CH), 8.00 – 7.95 (3H, m, Ar*H*), 7.93 – 7.87 (4H, m, Ar*H*), 7.82 – 7.75 (6H, m, Ar*H*), 7.67 (2H, m, Ar*H*), 7.53 – 7.45 (4H, m, Ar*H*, $\mathrm{CON}H_2$), 7.37 (1H, d, *J* = 8.6 Hz, Ar*H*), 4.90 (1H, t, *J* = 8.1 Hz, C*H*NH), 4.25 (1H, hept, *J* = 6.1 Hz, $\mathrm{C}H\mathrm{Me_2}$), 4.07 (1H, d, *J* = 8.1 Hz, C*H*OMe), 3.31 (3H, s, $\mathrm{OC}H_3$), 3.01 (2H, t, *J* = 7.5 Hz, $\mathrm{ArC}H_2\mathrm{CH_2}$), 2.71 (2H, t, *J* = 7.5 Hz, $\mathrm{ArCH_2C}H_2$), 1.02 (6H, d, *J* = 6.1 Hz, $\mathrm{CH(C}H_3\mathrm{)_2}$) ppm; ^{13}C-NMR (100 MHz, DMSO-d_6): δ = 171.0 (q, $\mathrm{CONH_2}$), 170.4 (q, *C*ONH), 168.2 (q, *C*ONH), 167.5 (q, *C*ONH), 166.8 (q, $C\mathrm{O_2H}$), 165.4 (q, *C*ONH), 154.5 (q, C-Ar), 147.3 (q, C-Ar), 146.3 (q, $\mathrm{C\text{-}Ar_{Triazol}}$), 142.1 (q, C-Ar), 141.9 (q, C-Ar), 138.9 (q, C-Ar), 138.7 (q, C-Ar), 134.4 (q, C-Ar), 132.2 (2C, t, C-Ar), 130.3 (2C, t, C-Ar), 129.5 (2C, t, C-Ar), 128.3 (2C, t, C-Ar), 128.0 (q, C-Ar), 126.5 (q, C-Ar), 125.8 (2C, t, C-Ar), 125.4 (q, C-Ar), 123.5 (t, C-Ar), 122.6 (t, $\mathrm{C\text{-}Ar_{Triazol}}$), 120.6 (2C, t, C-Ar), 119.8 (2C, t, C-Ar), 119.0 (q, C-Ar), 118.4 (q, CN), 118.2 (2C, t, C-Ar), 114.8 (t, C-Ar), 108.9 (q, C-Ar), 80.1 (t, *C*HOMe), 75.7 (t, $C\mathrm{H(CH_3)_2}$), 57.7 (p, $\mathrm{O}C\mathrm{H_3}$), 55.6 (t, *C*HNH), 37.1 (s, $\mathrm{ArCH_2}C\mathrm{H_2}$), 30.6 (s, $\mathrm{Ar}C\mathrm{H_2CH_2}$), 21.8 (2C, p, $\mathrm{CH(}C\mathrm{H_3)_2}$) ppm; HRMS (ESI): *m/z* calc. for $\mathrm{C_{47}H_{43}N_9O_{10}}$ $[\mathrm{M + H}]^+$: 916.3031; found 916.3029.

4. Copies of ^{1}H- and ^{13}C-NMR spectra



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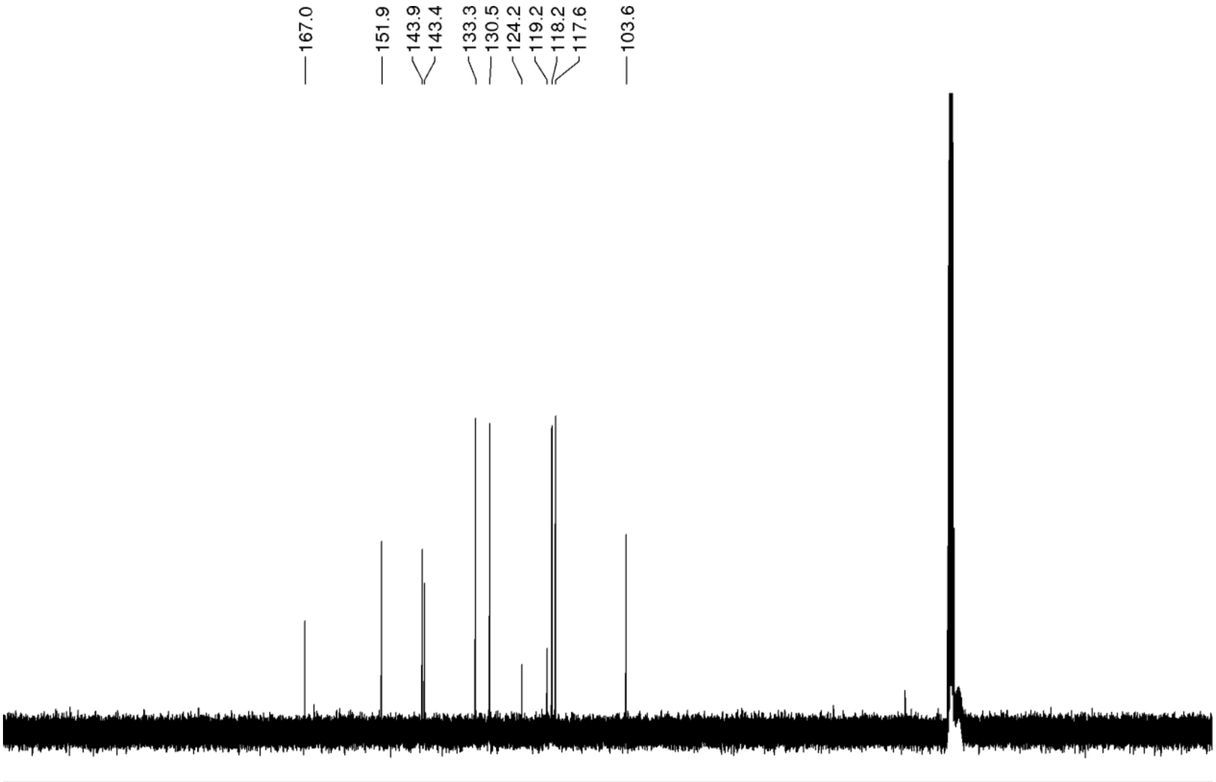

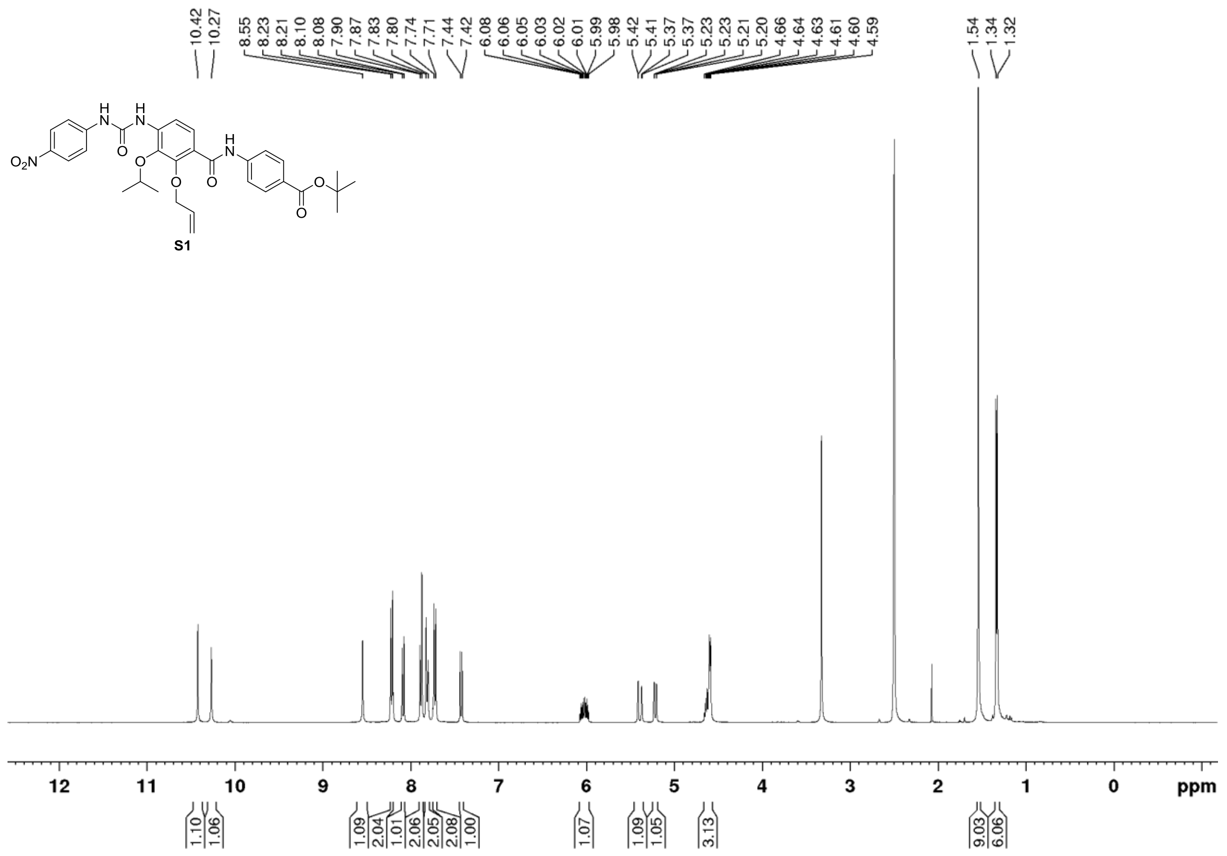
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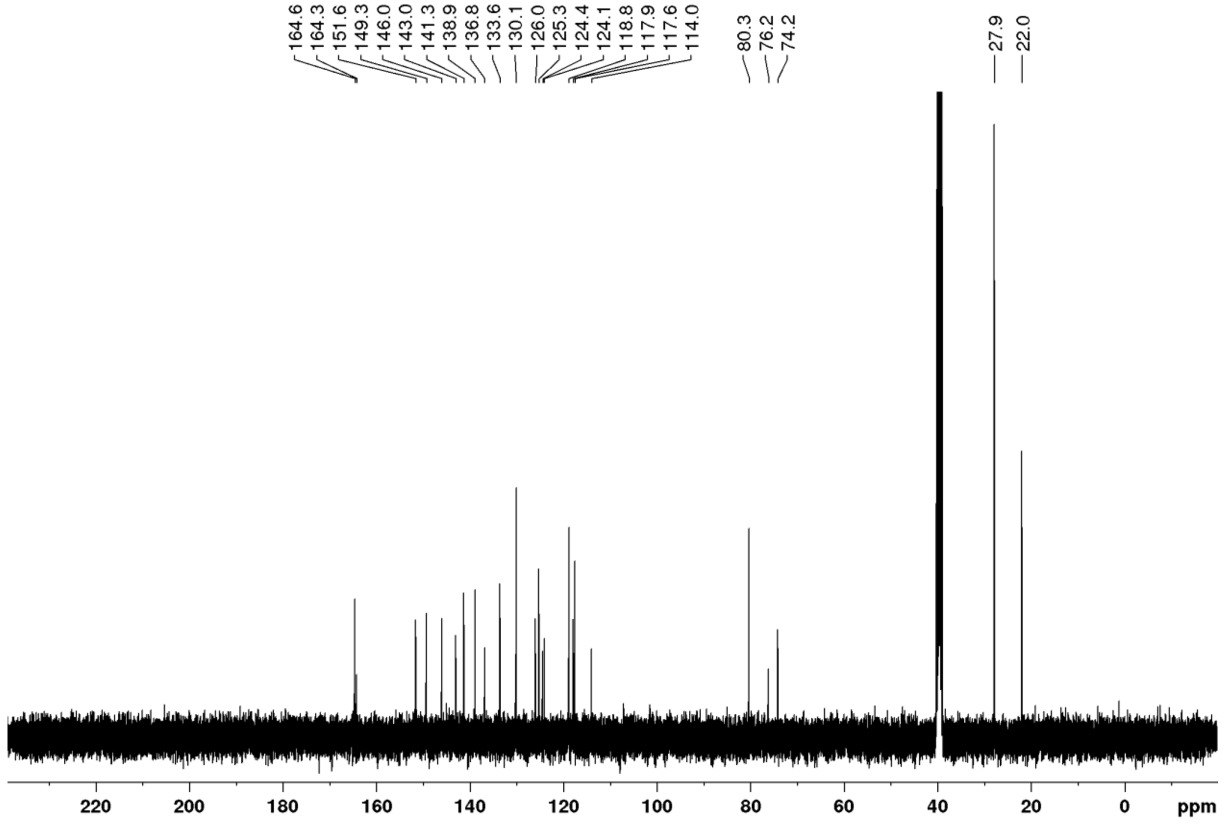
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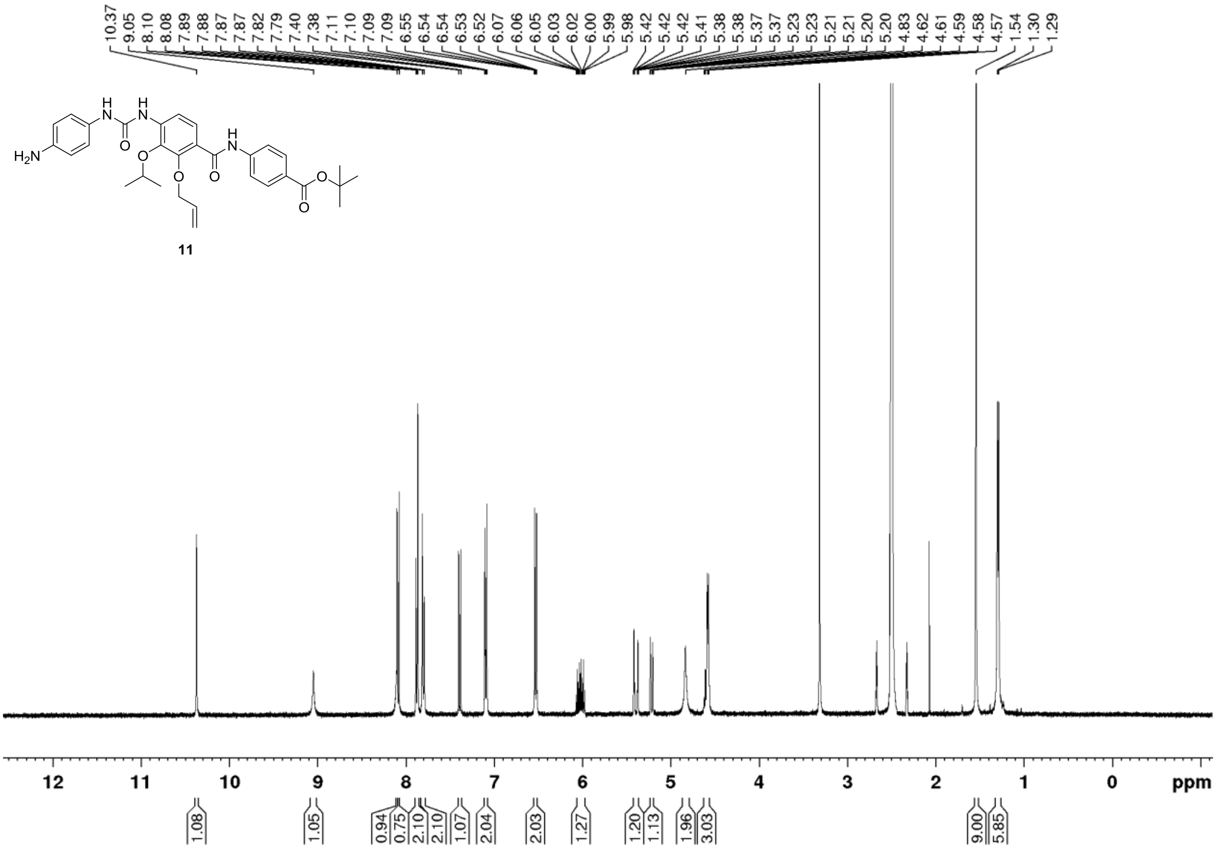
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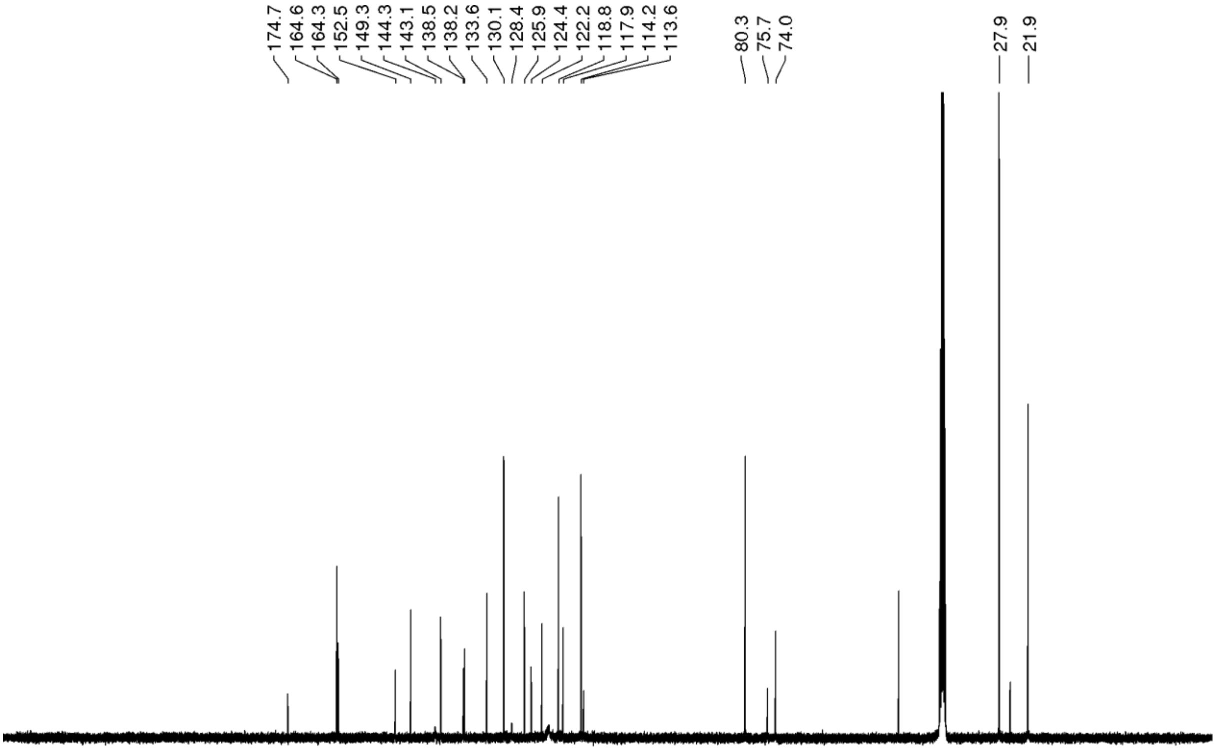
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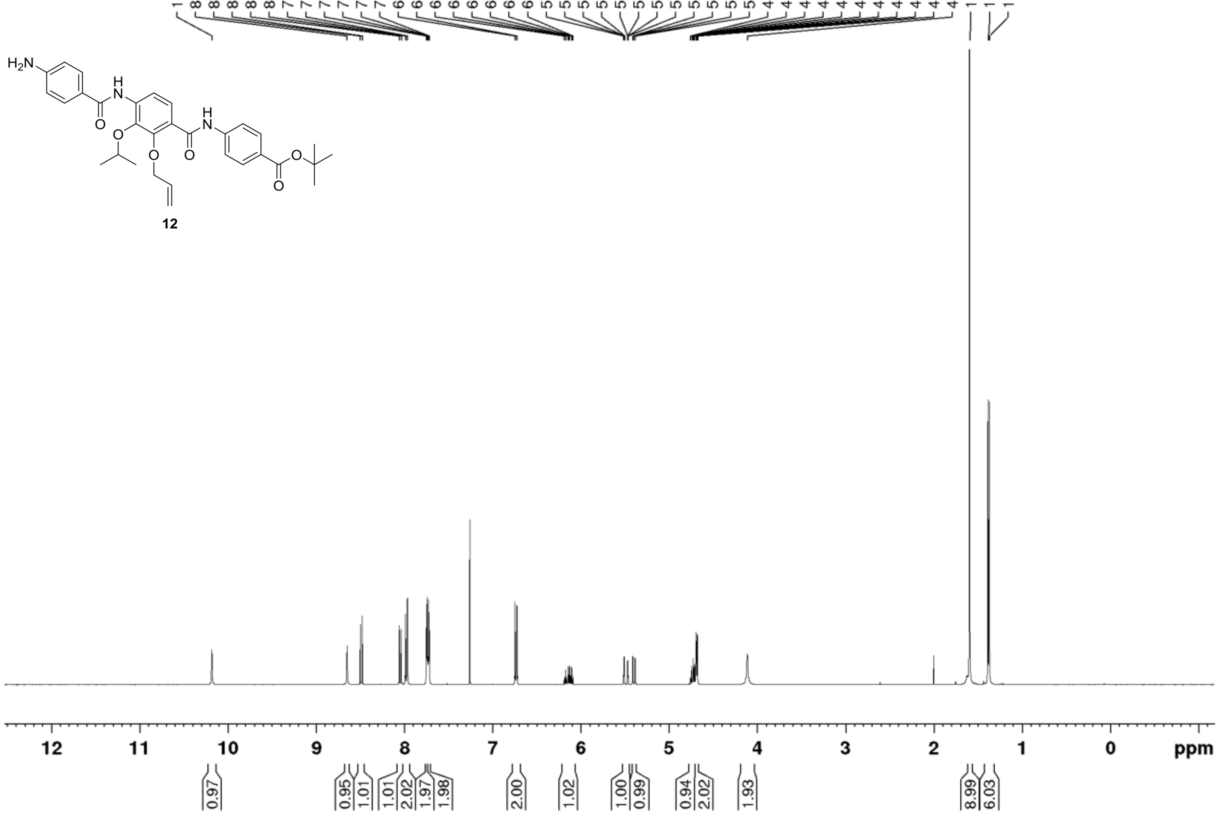
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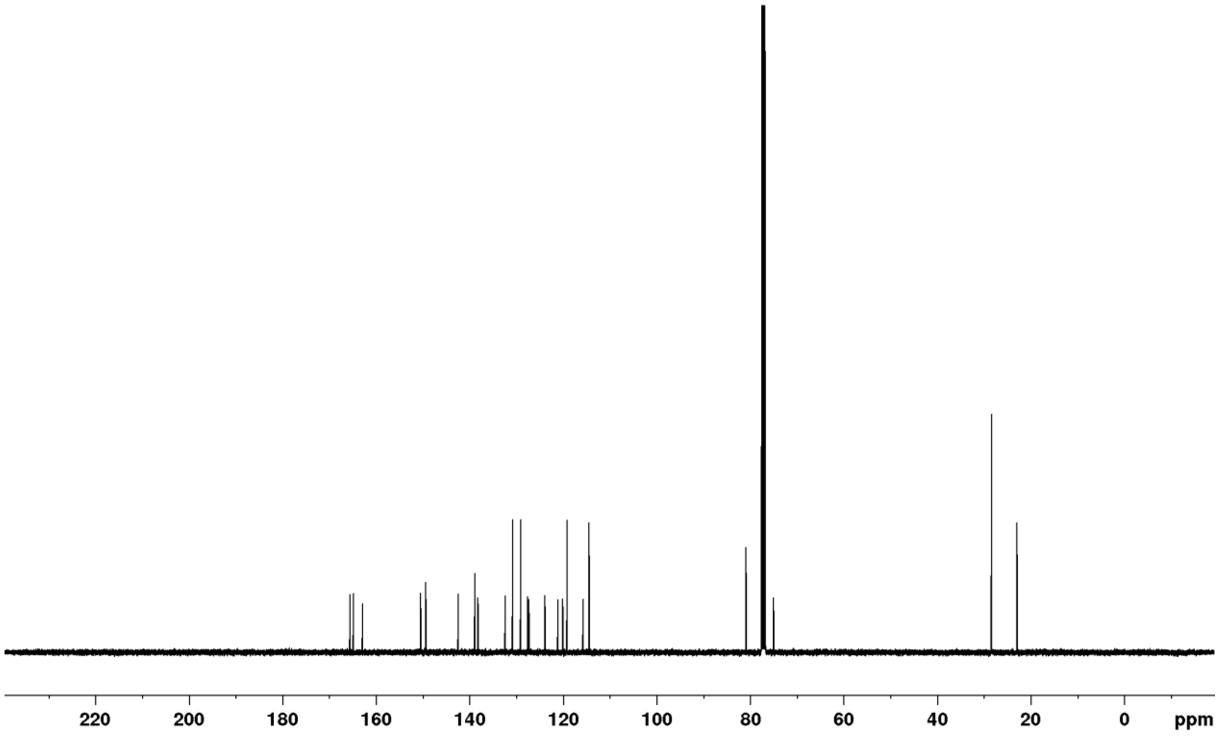
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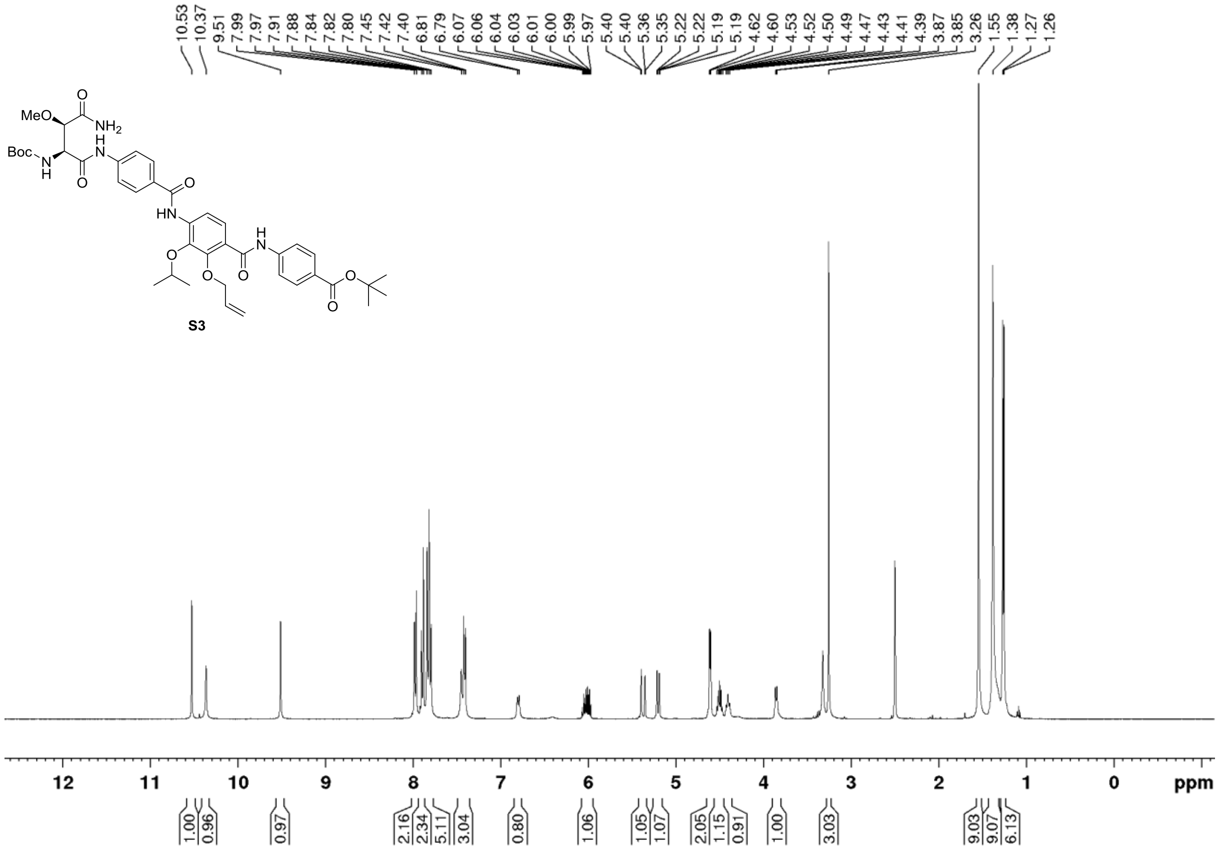
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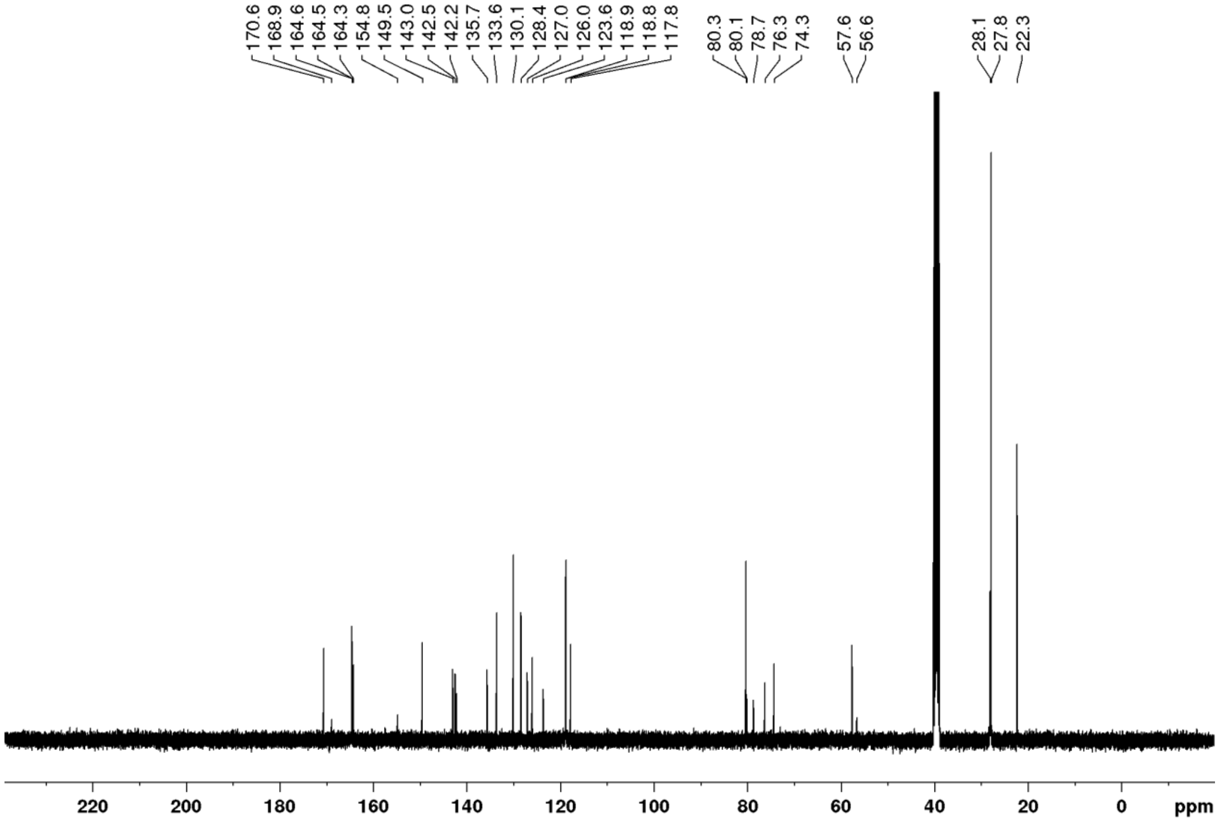
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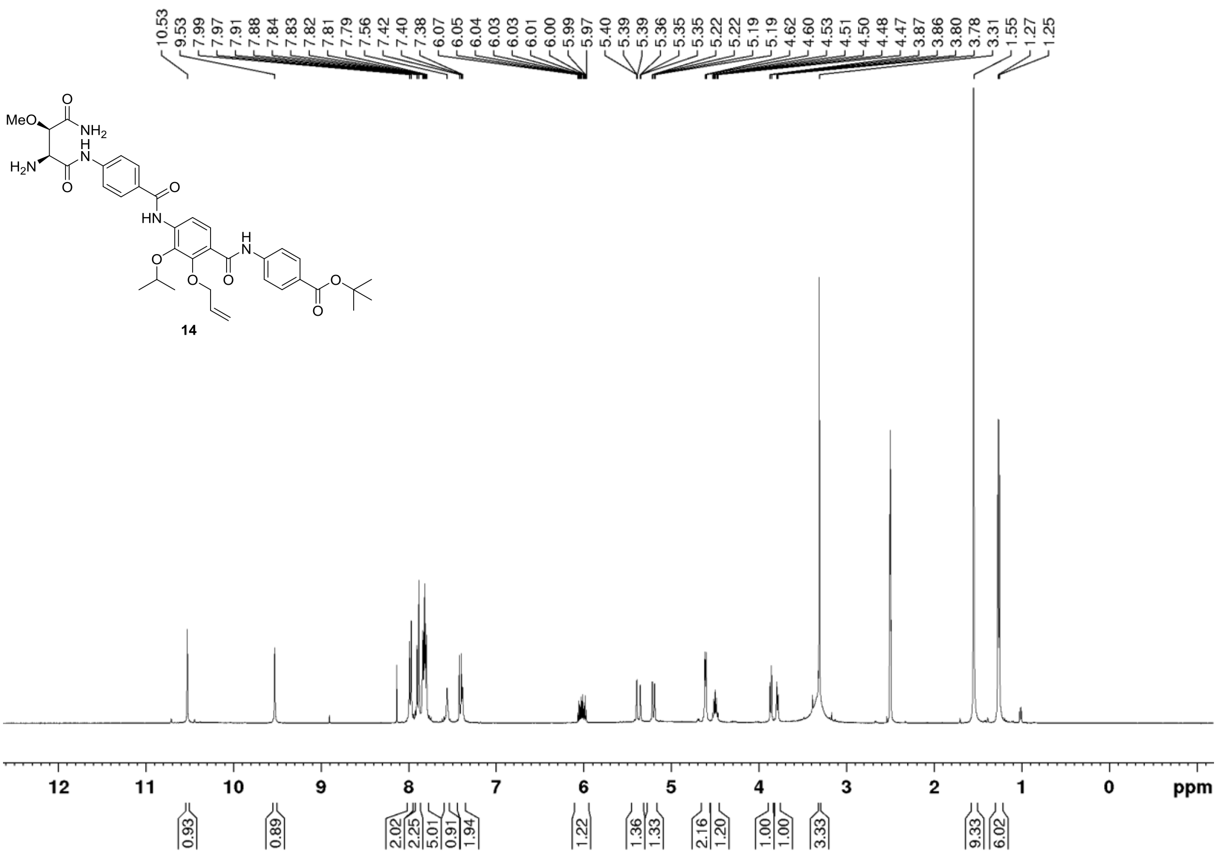
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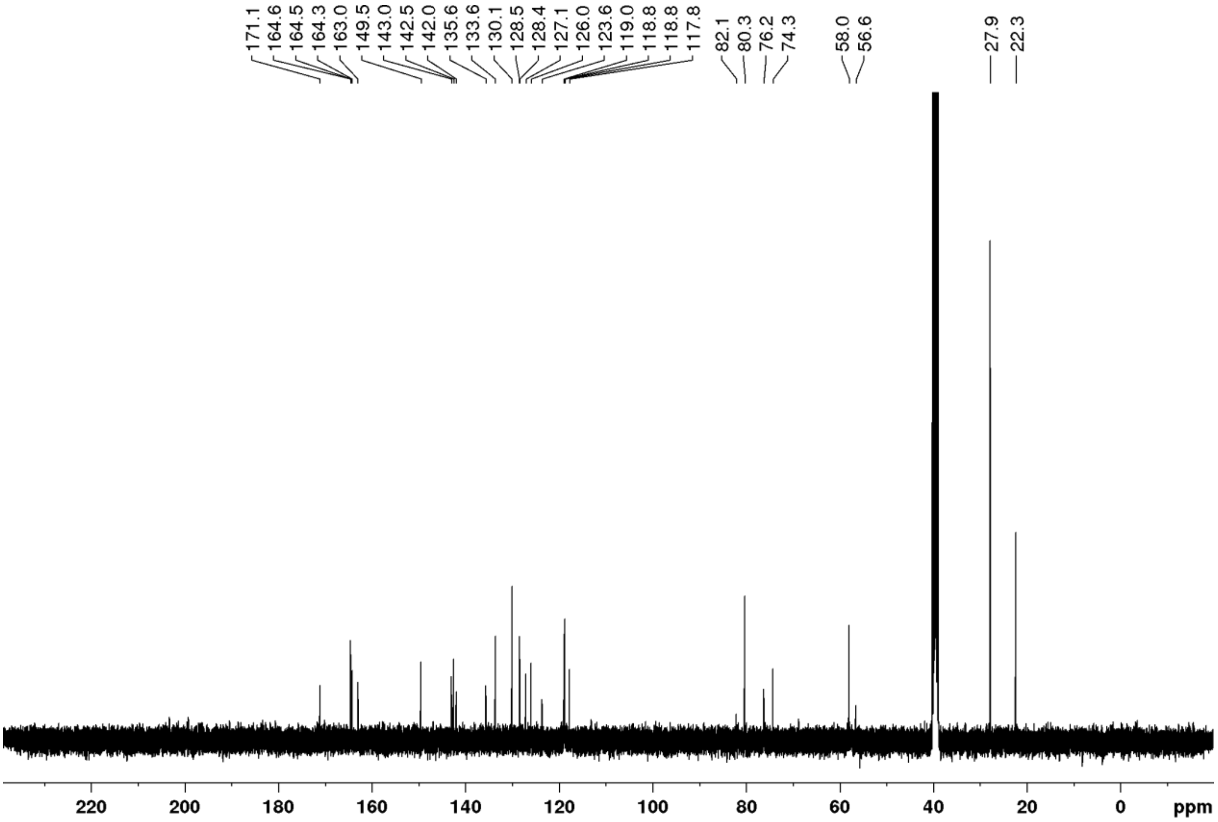
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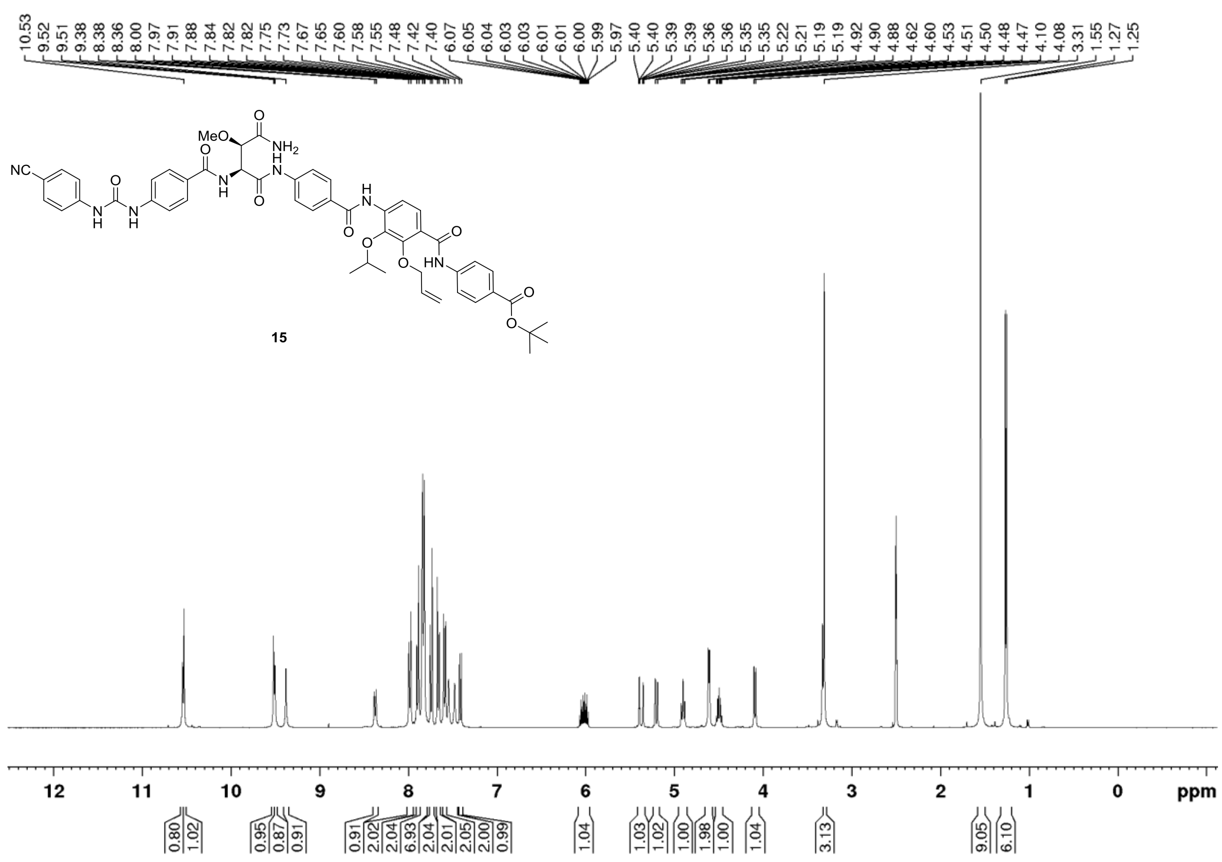
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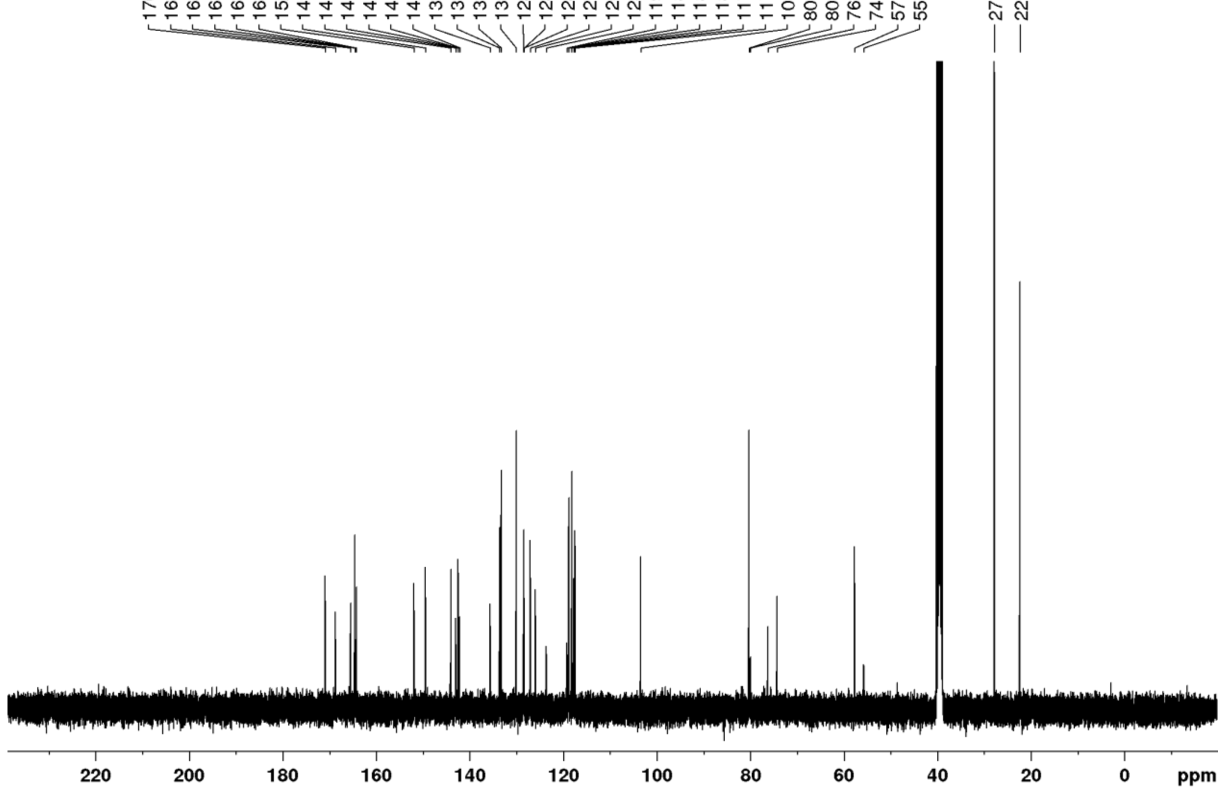
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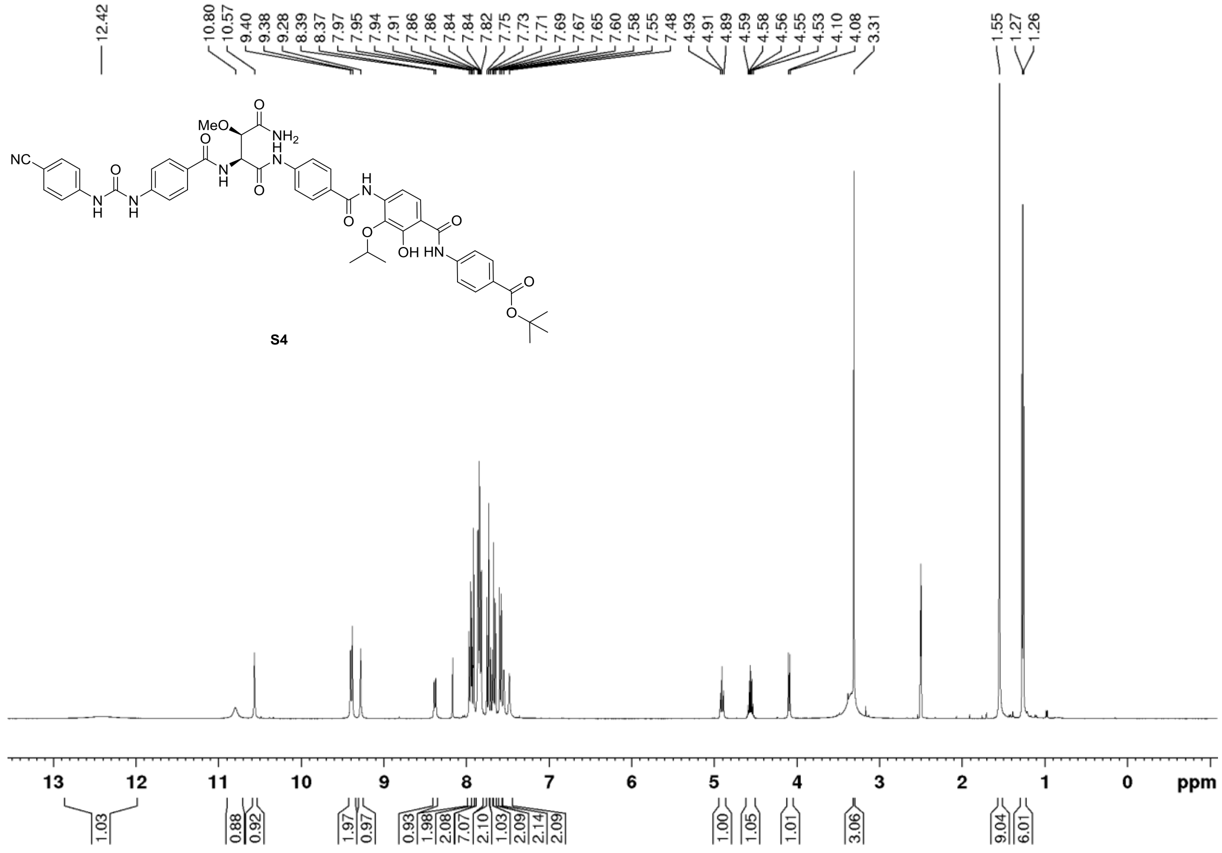
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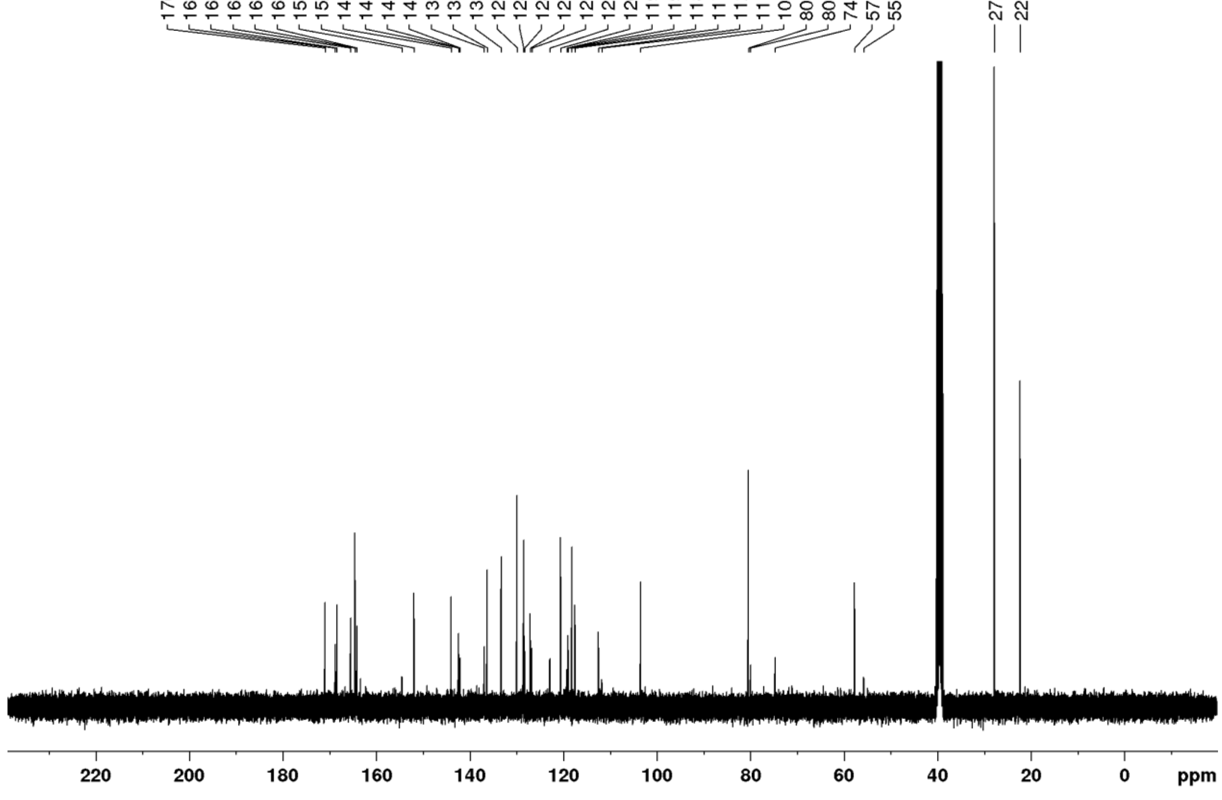
S4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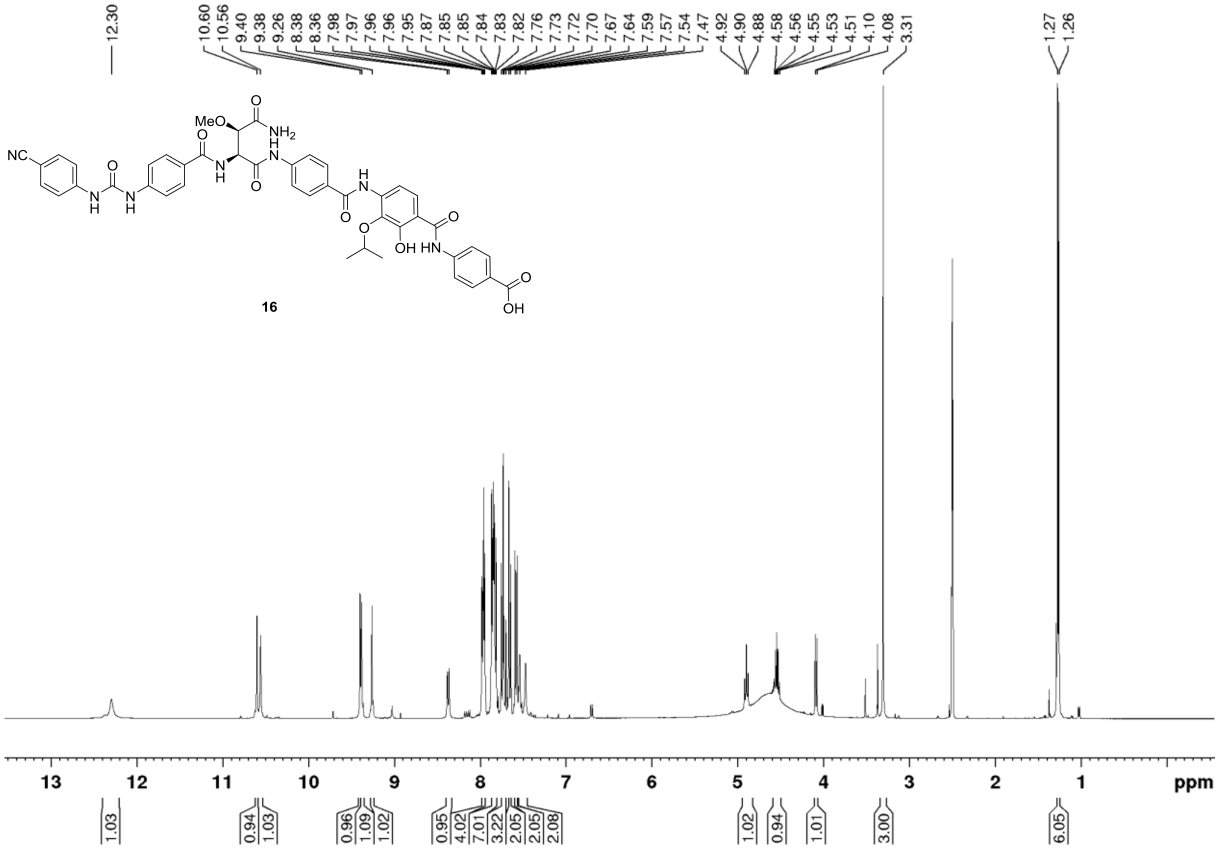
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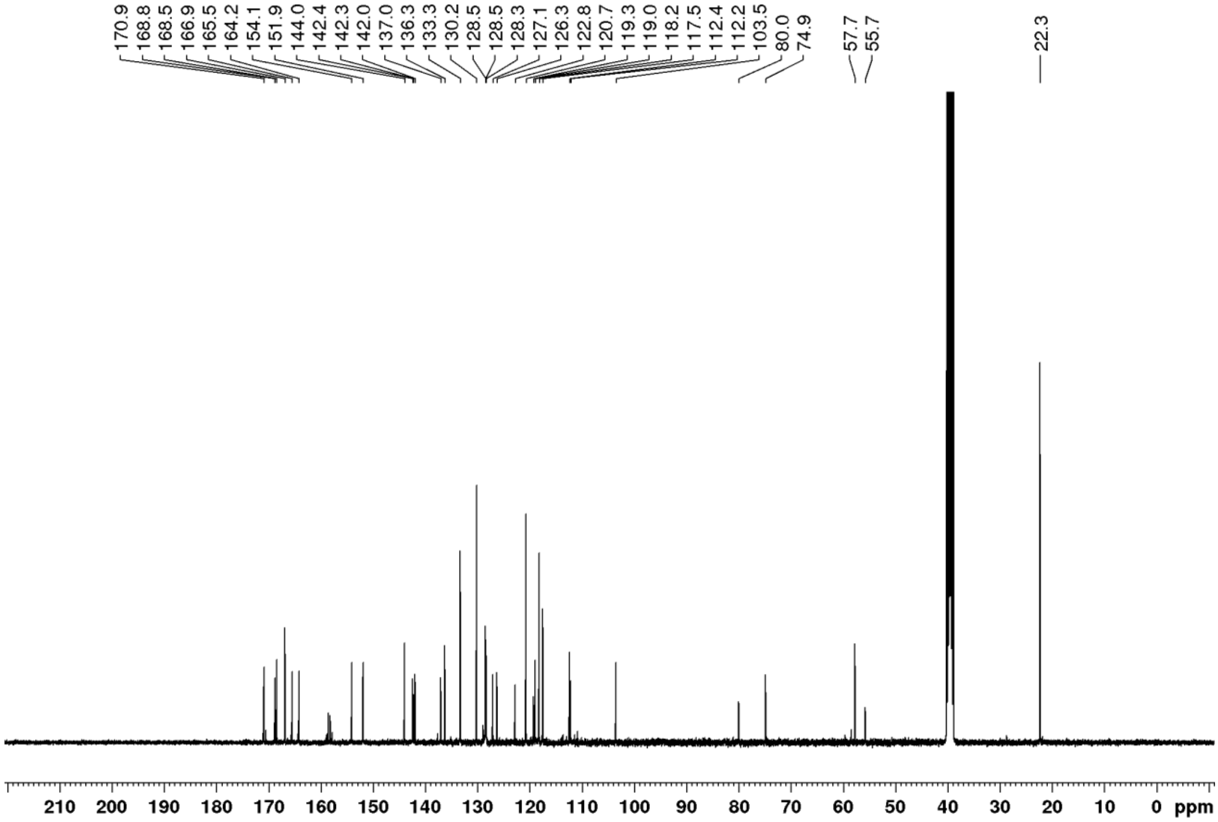
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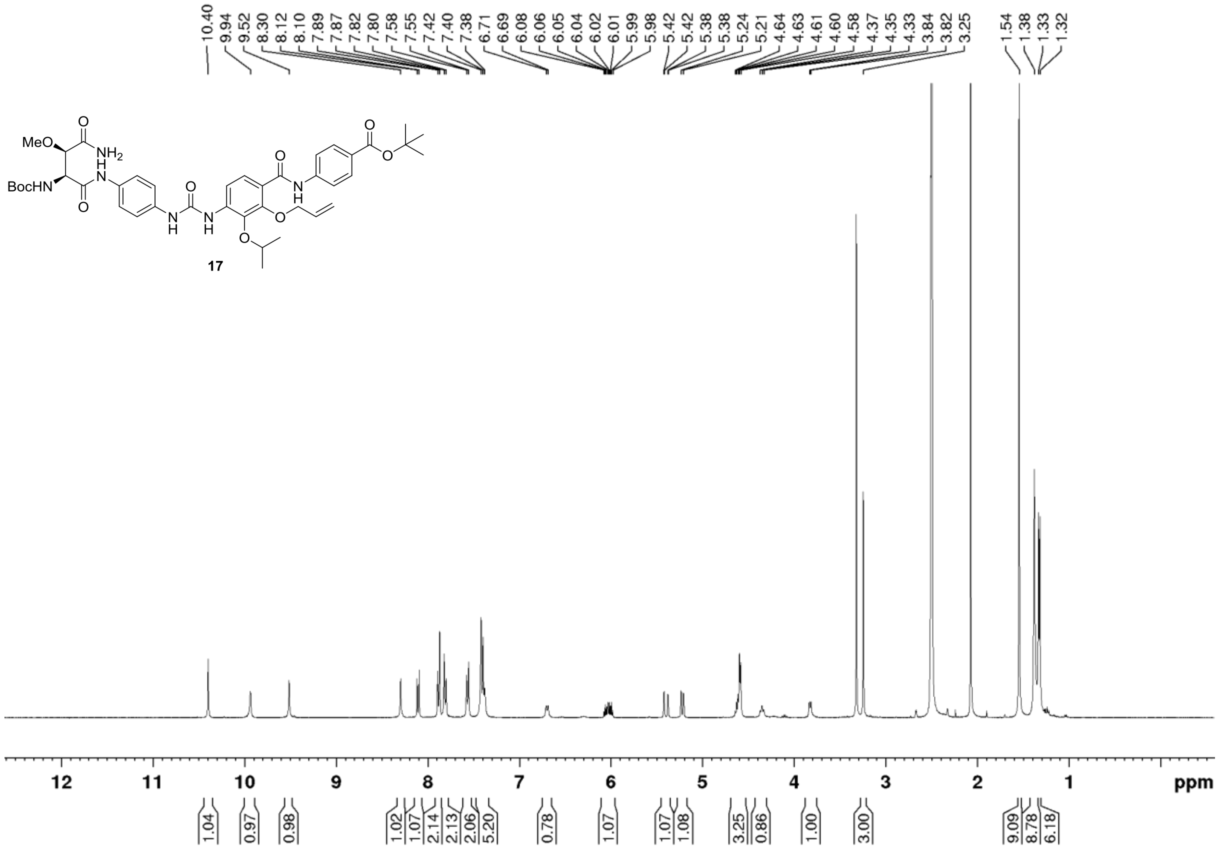
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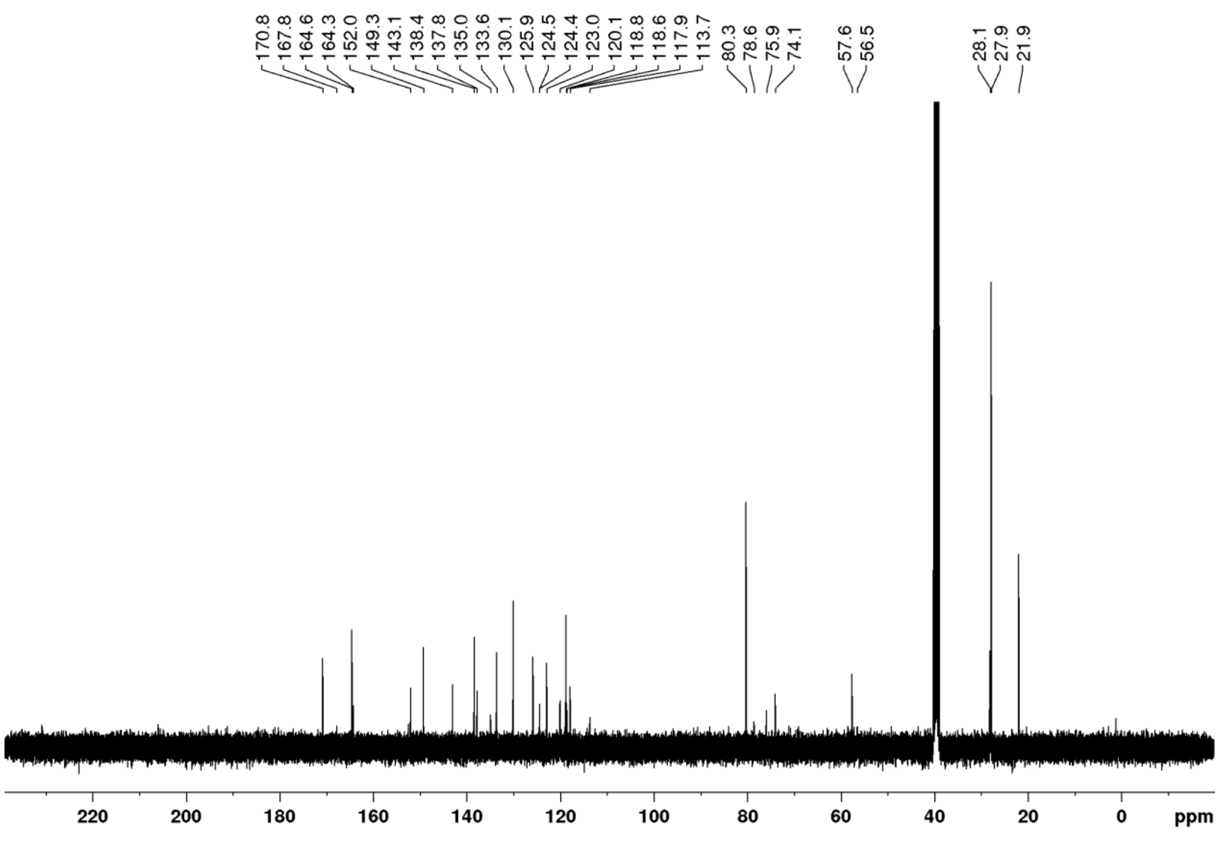
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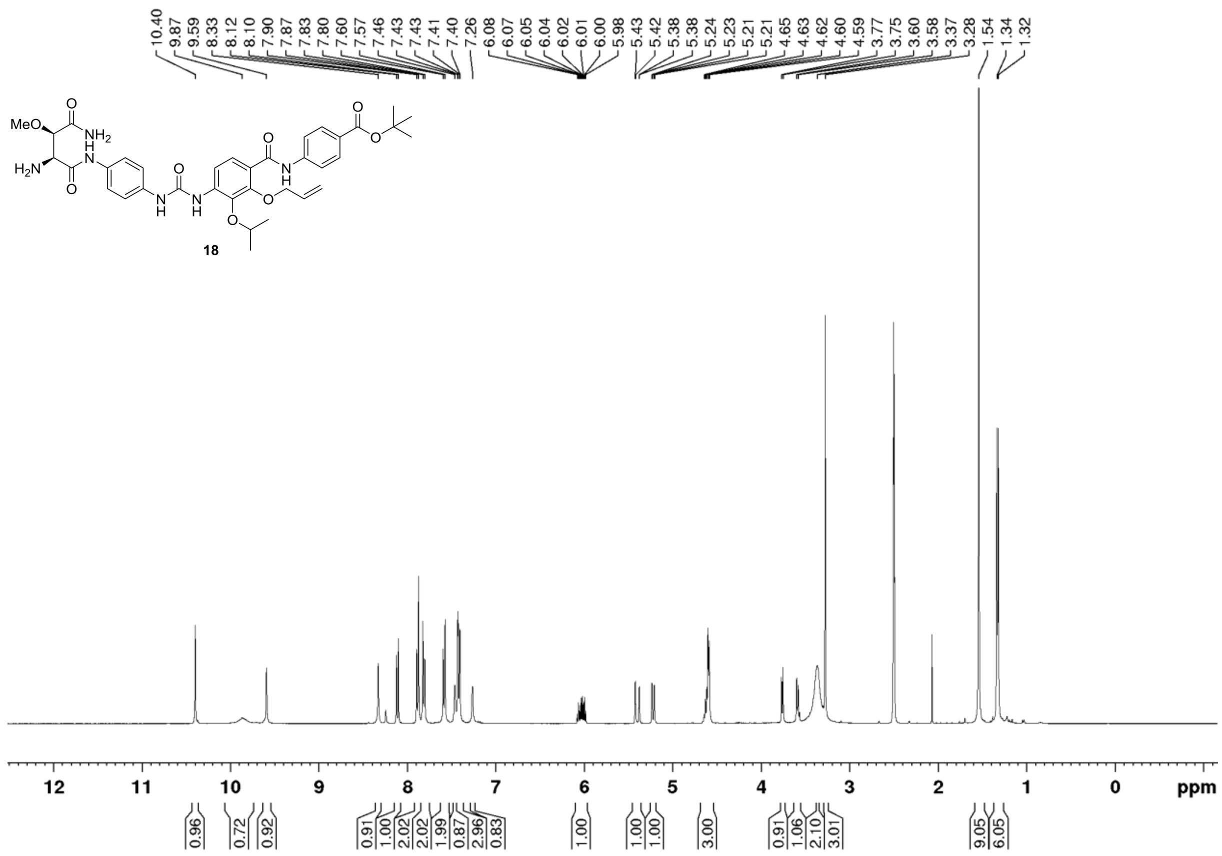

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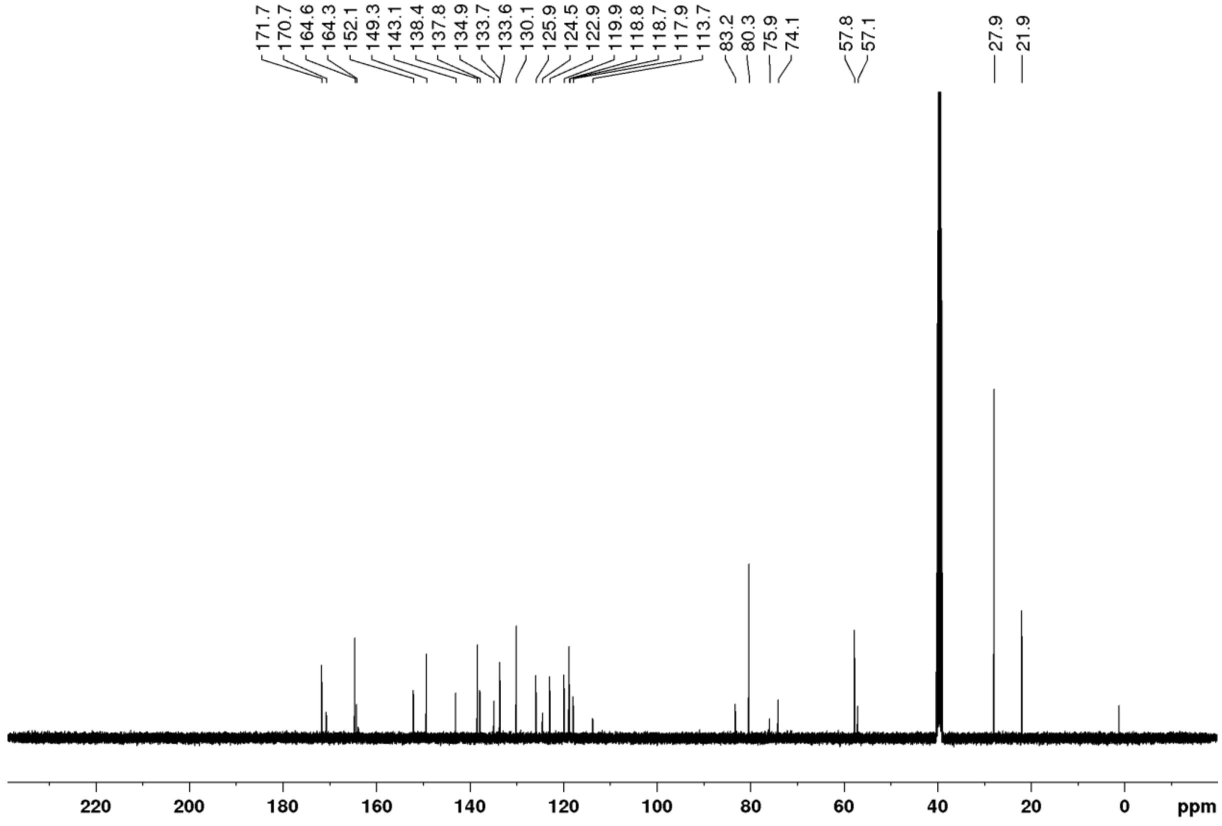

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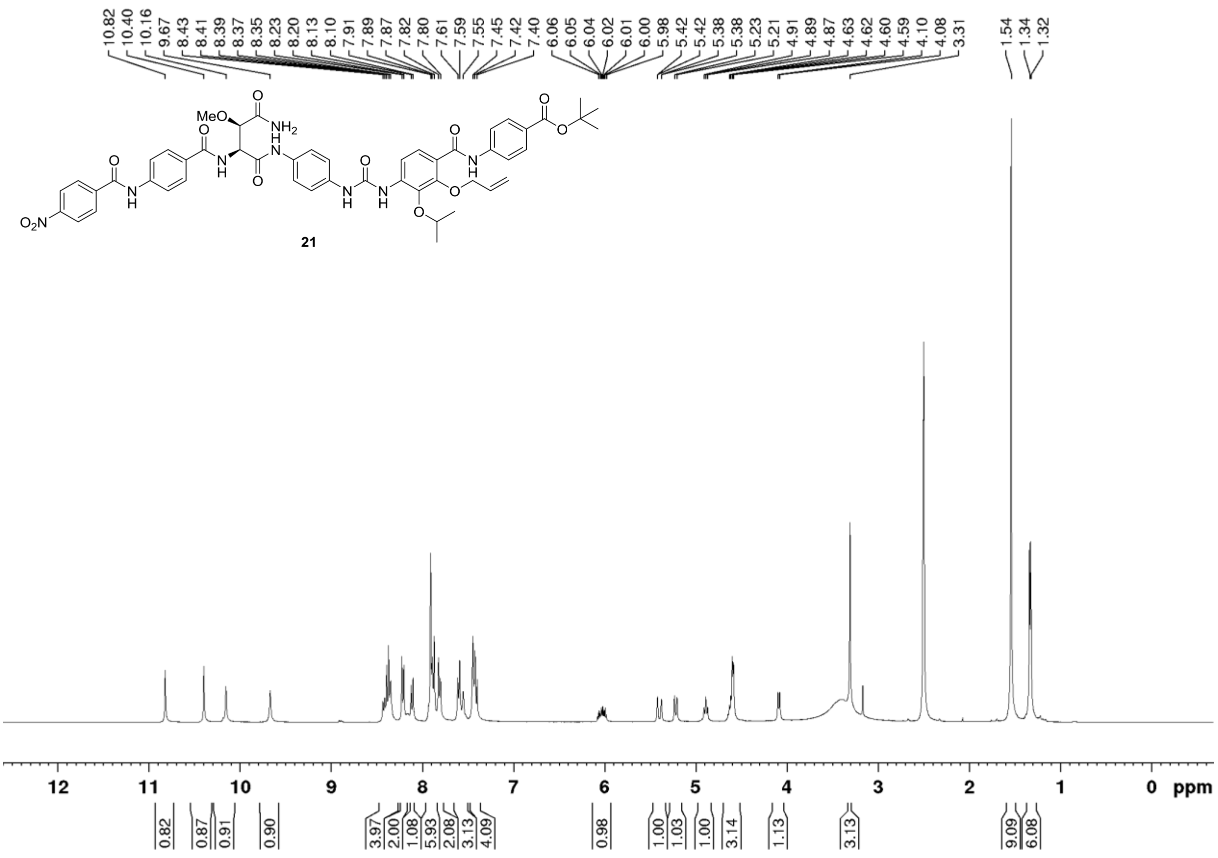
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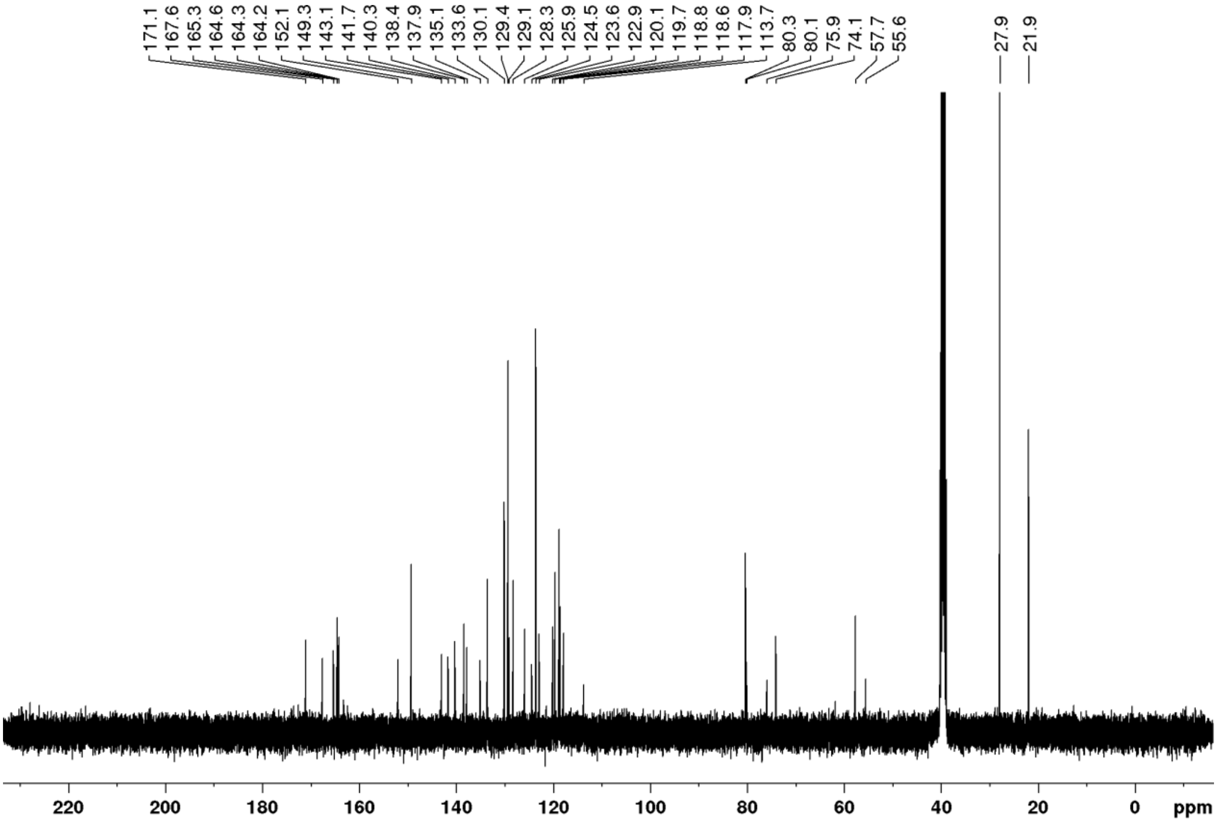
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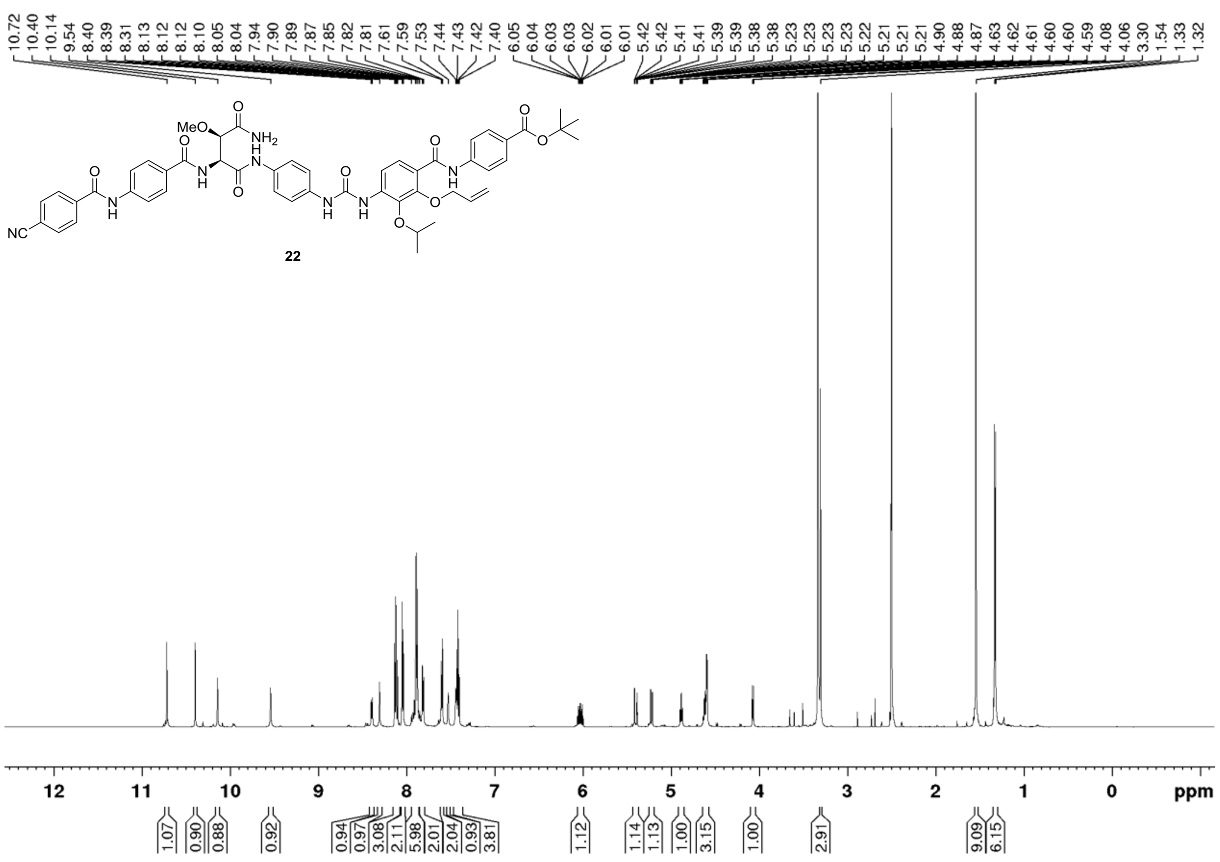
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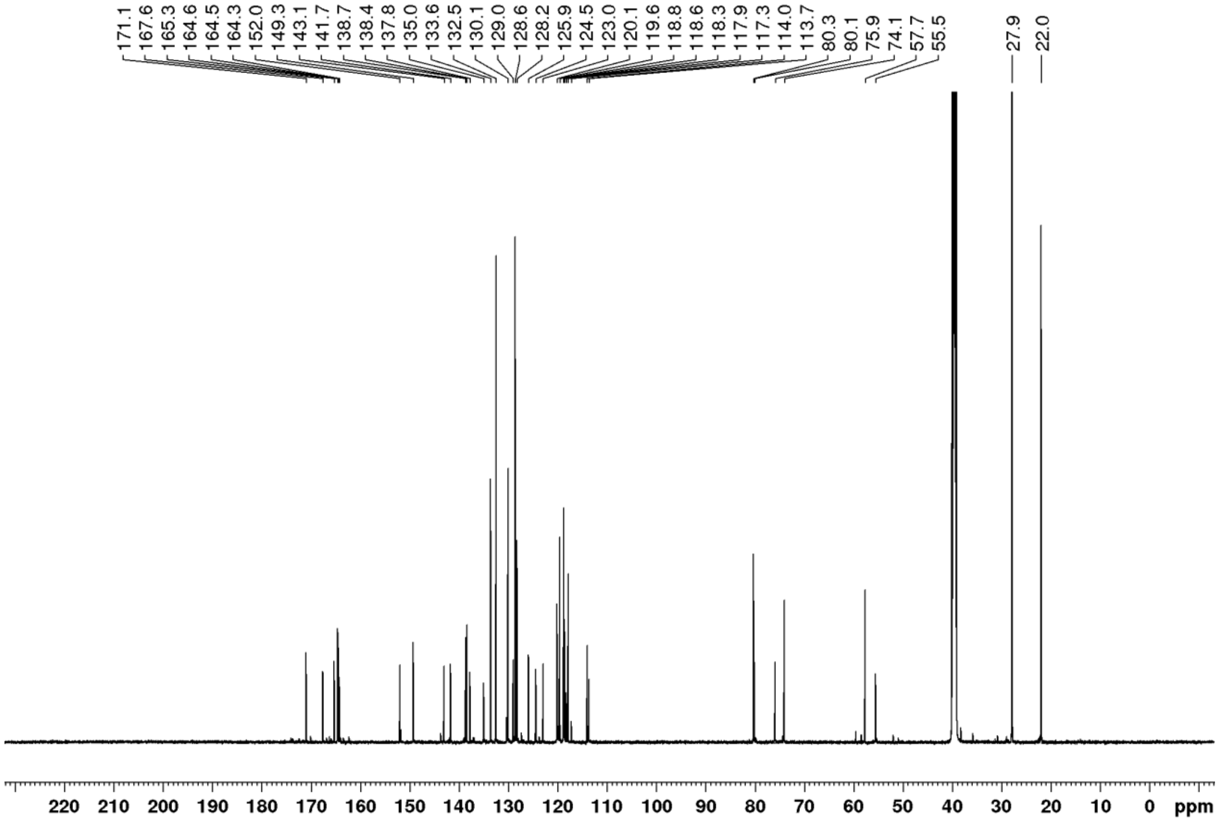
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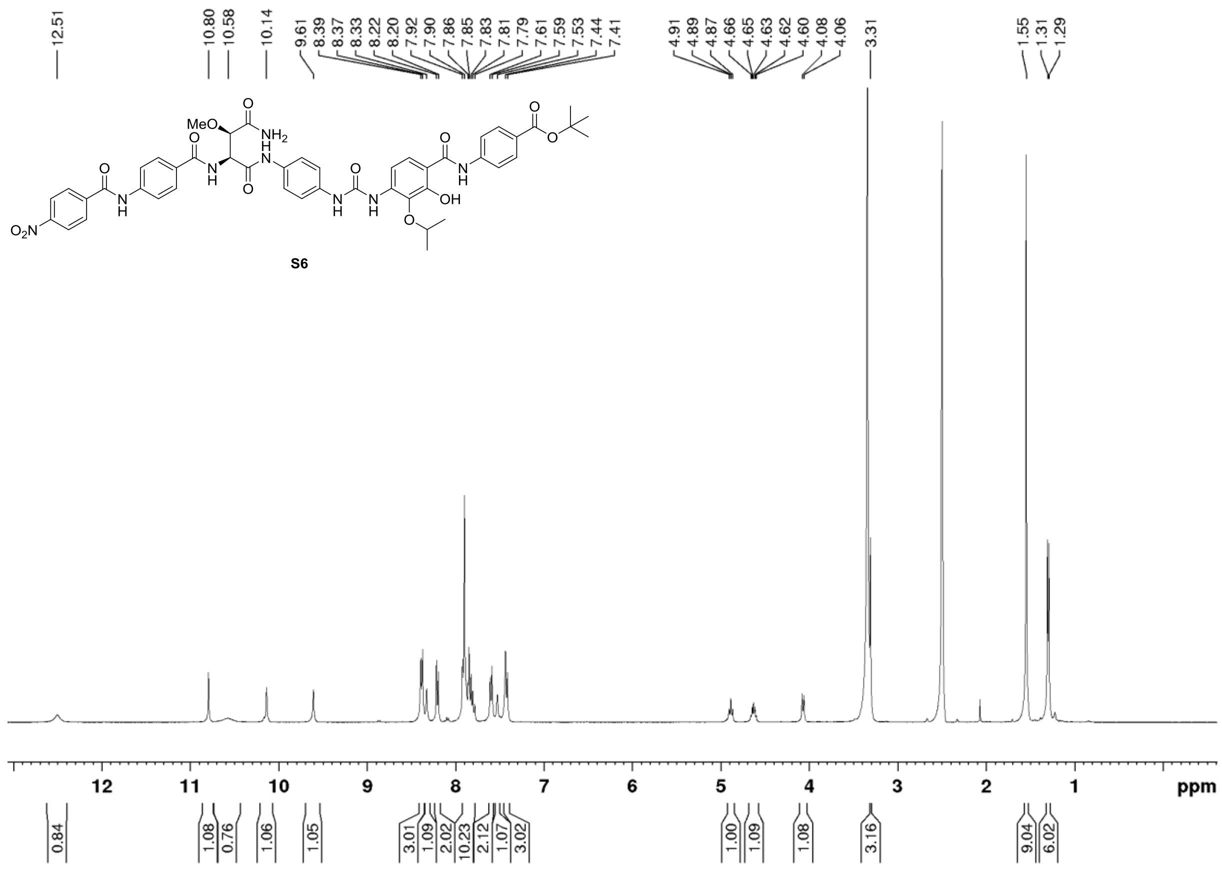
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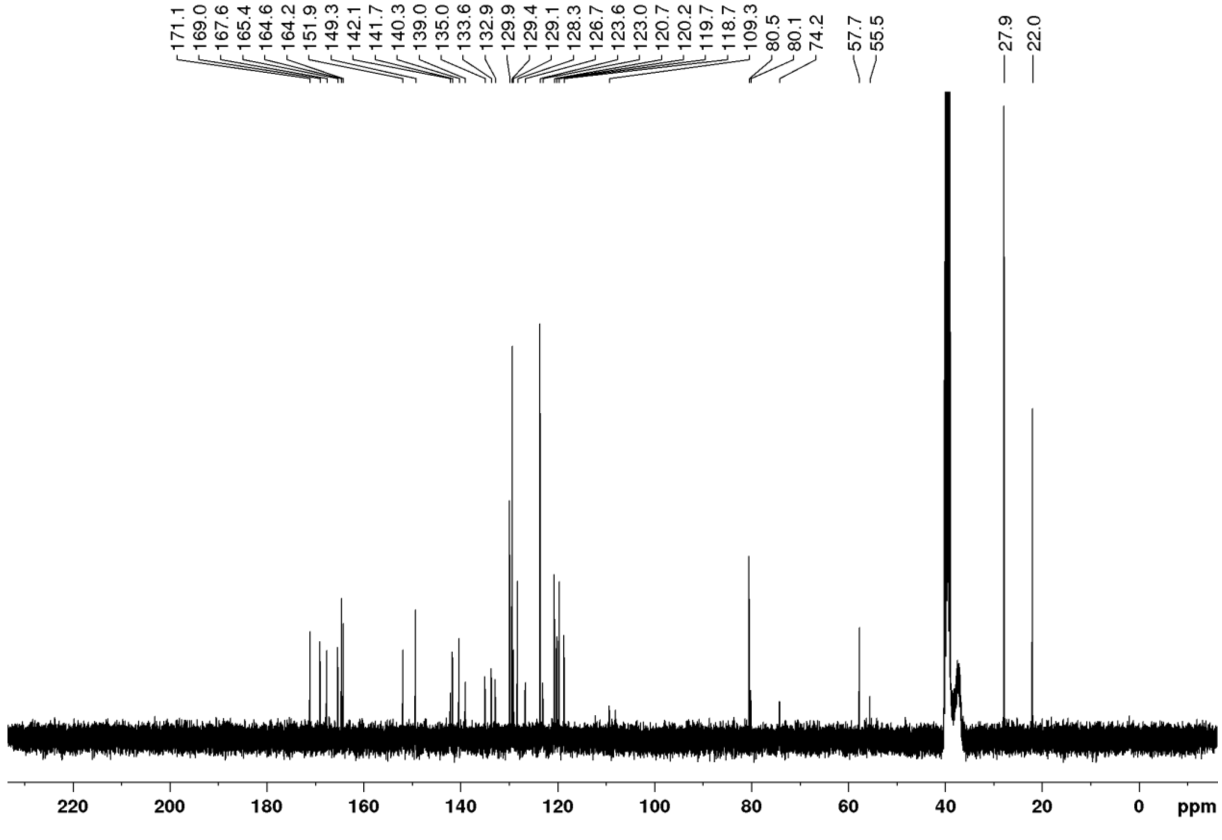
S6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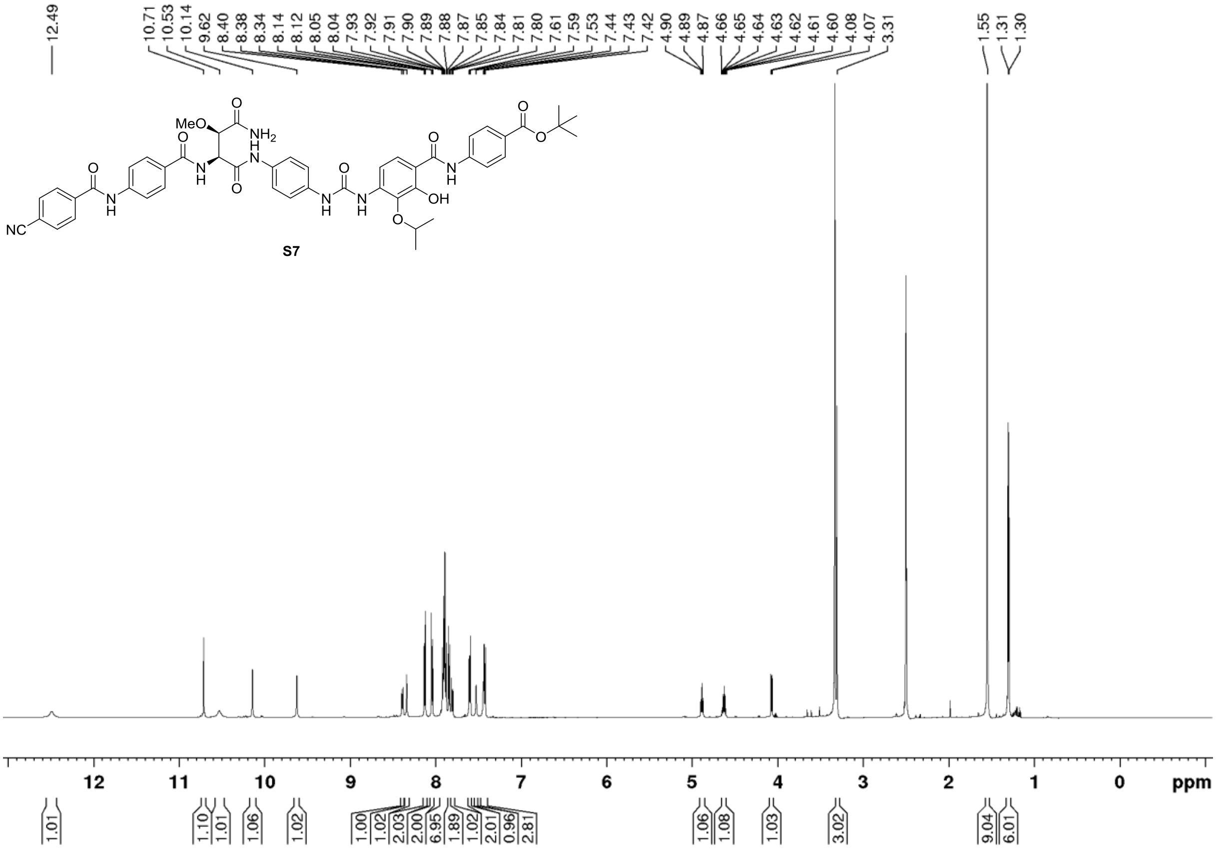
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N H
N H
N H
O
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O
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OH
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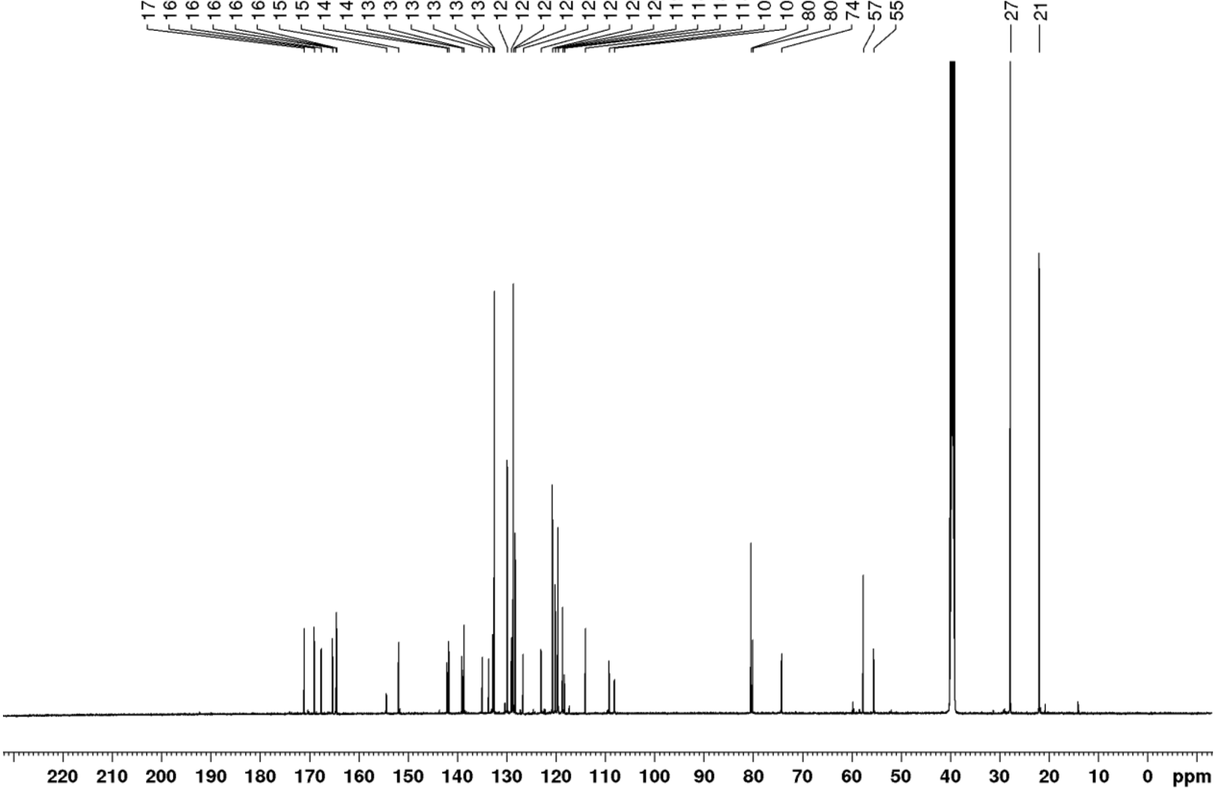
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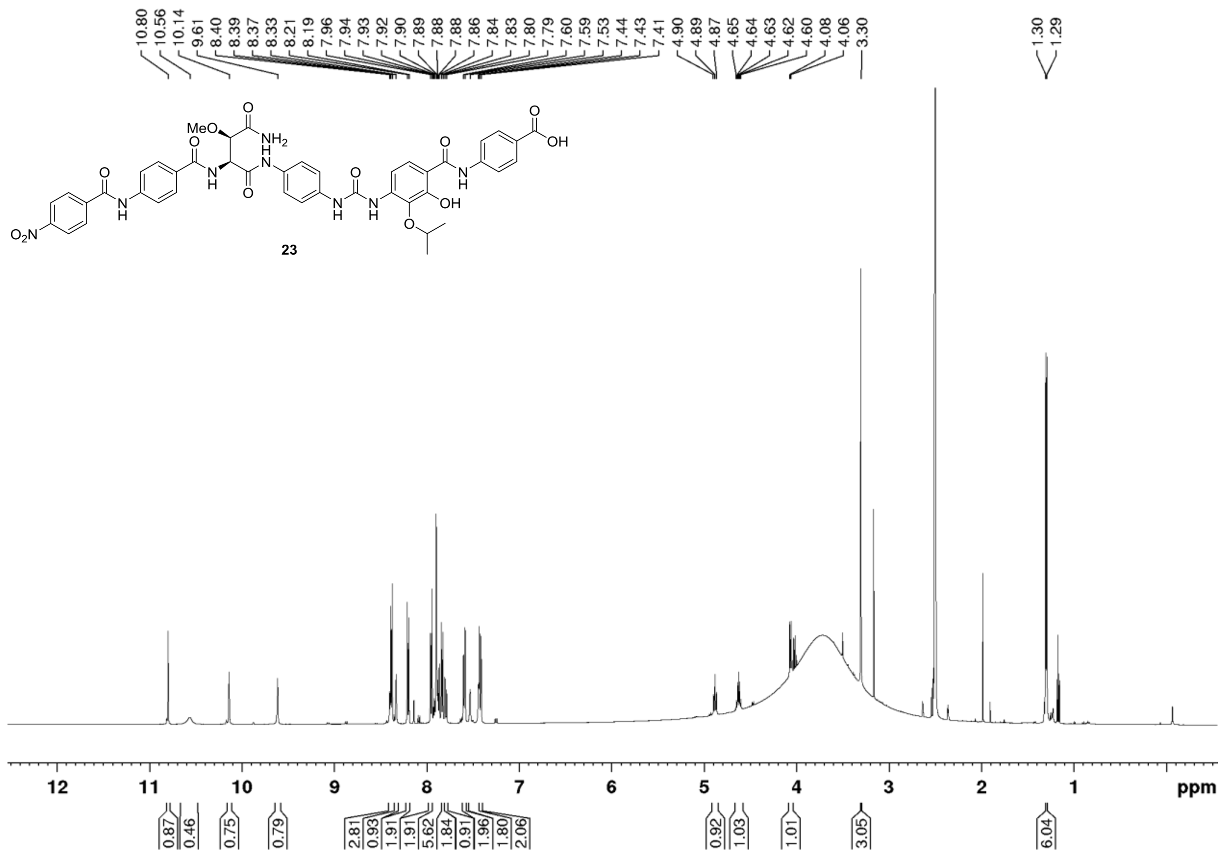
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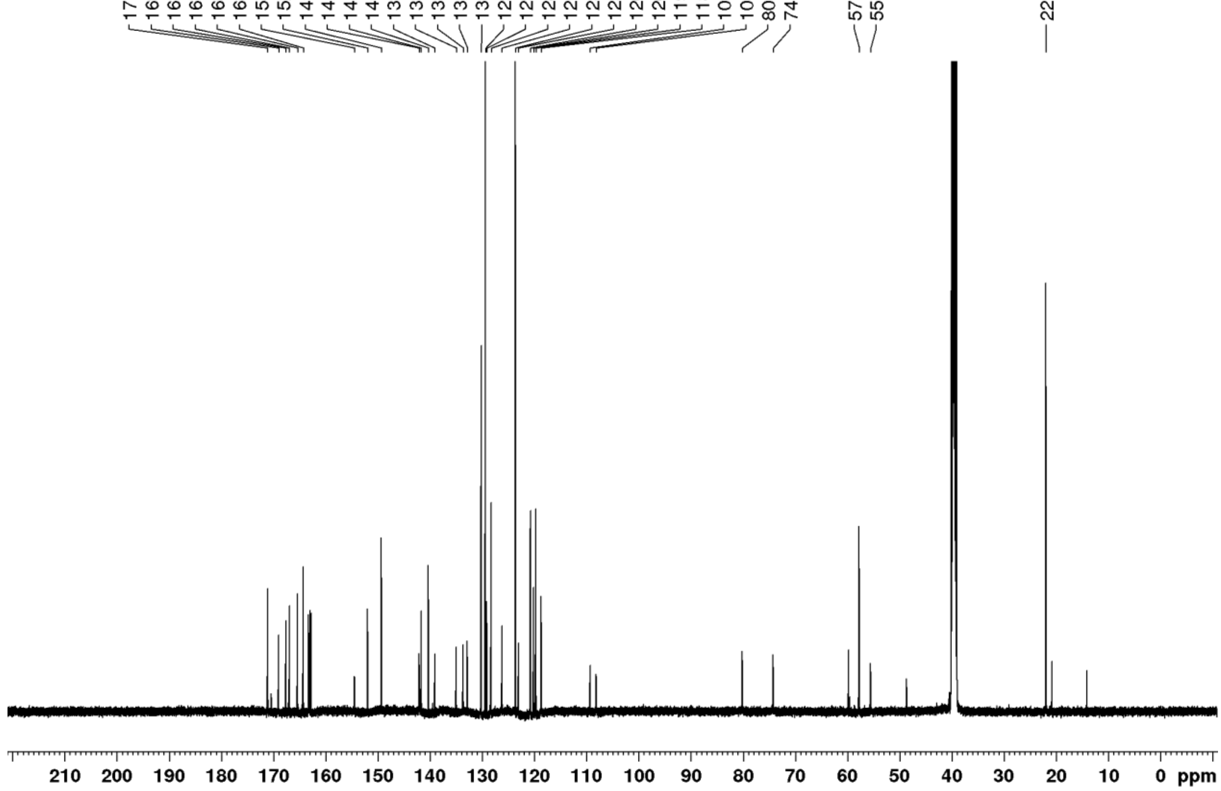
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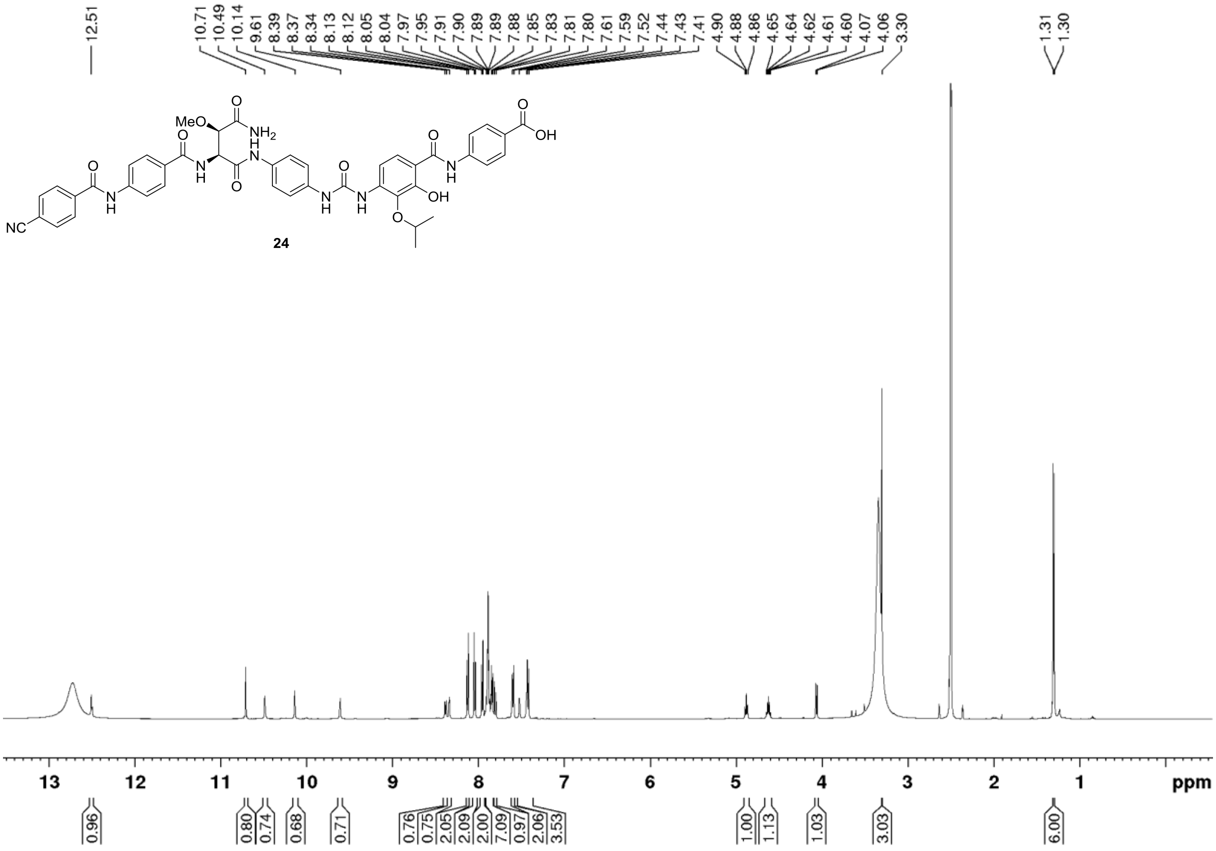
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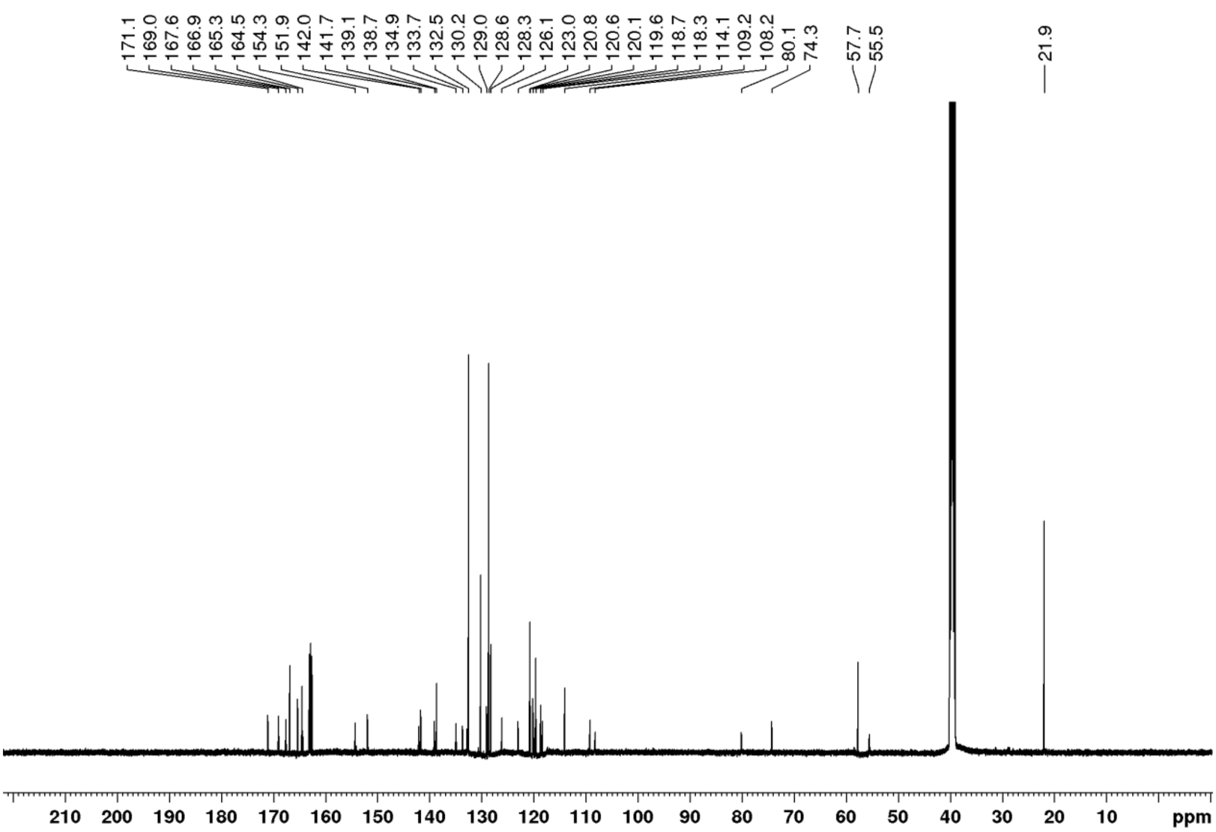
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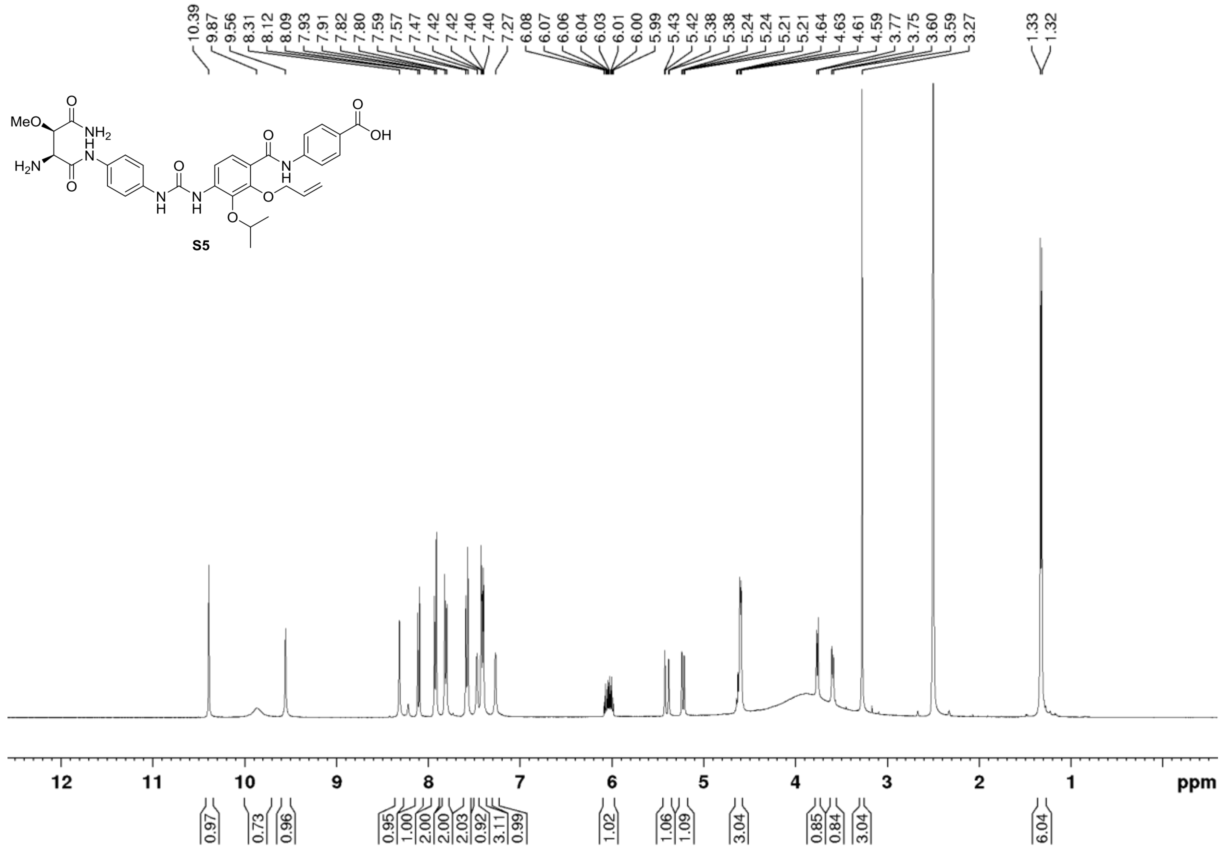
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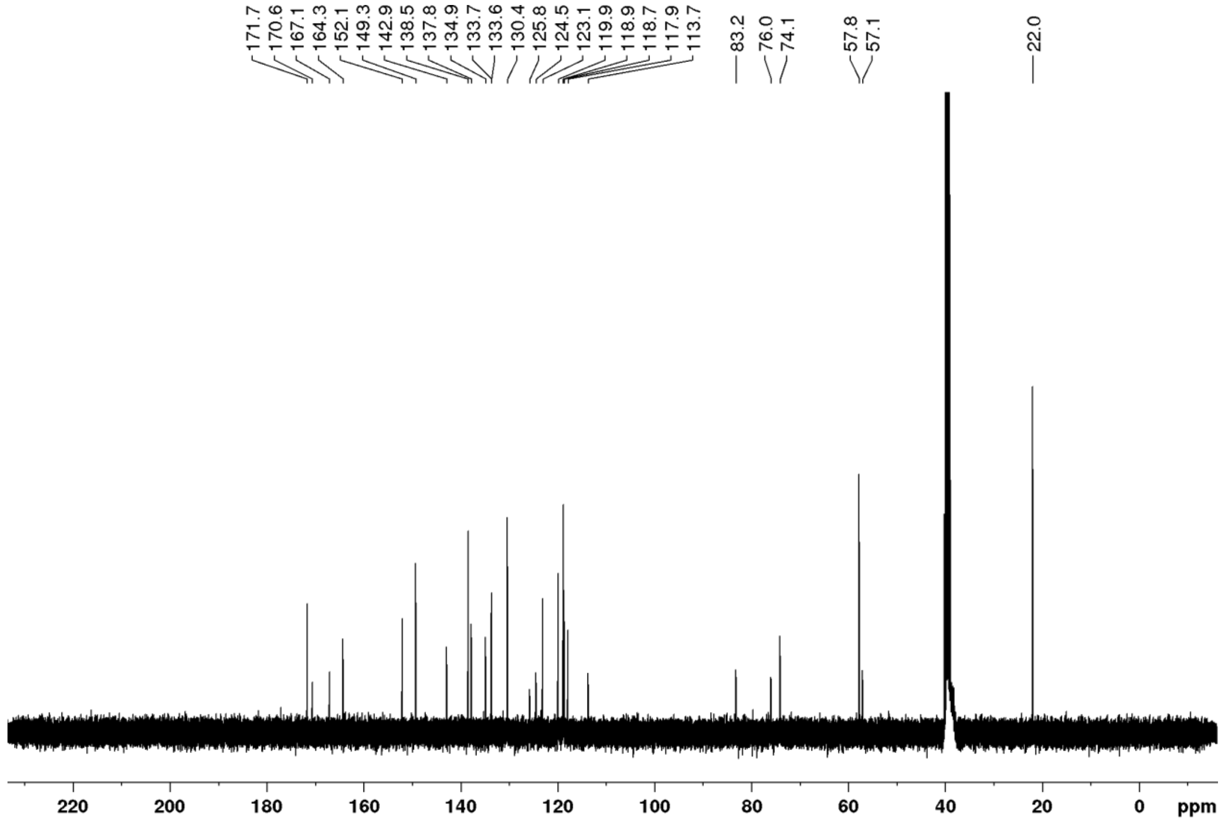
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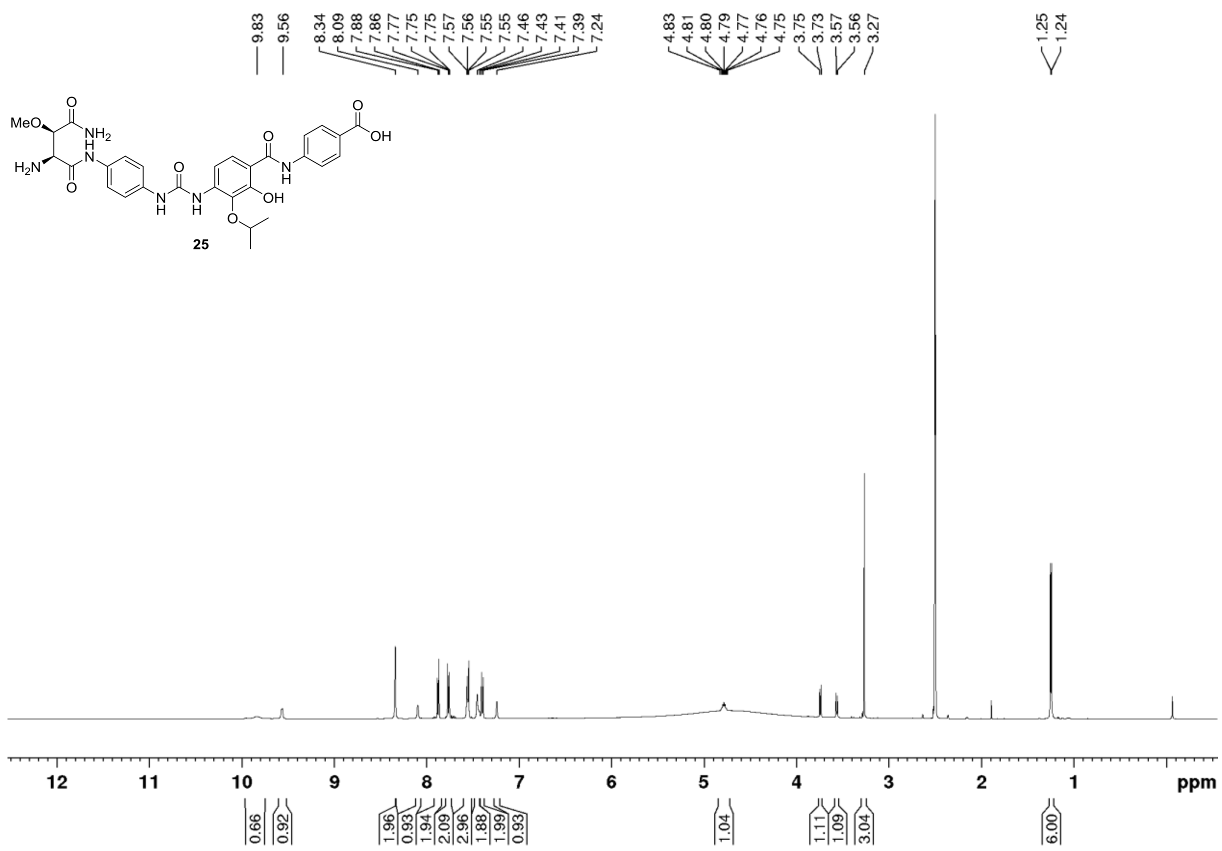
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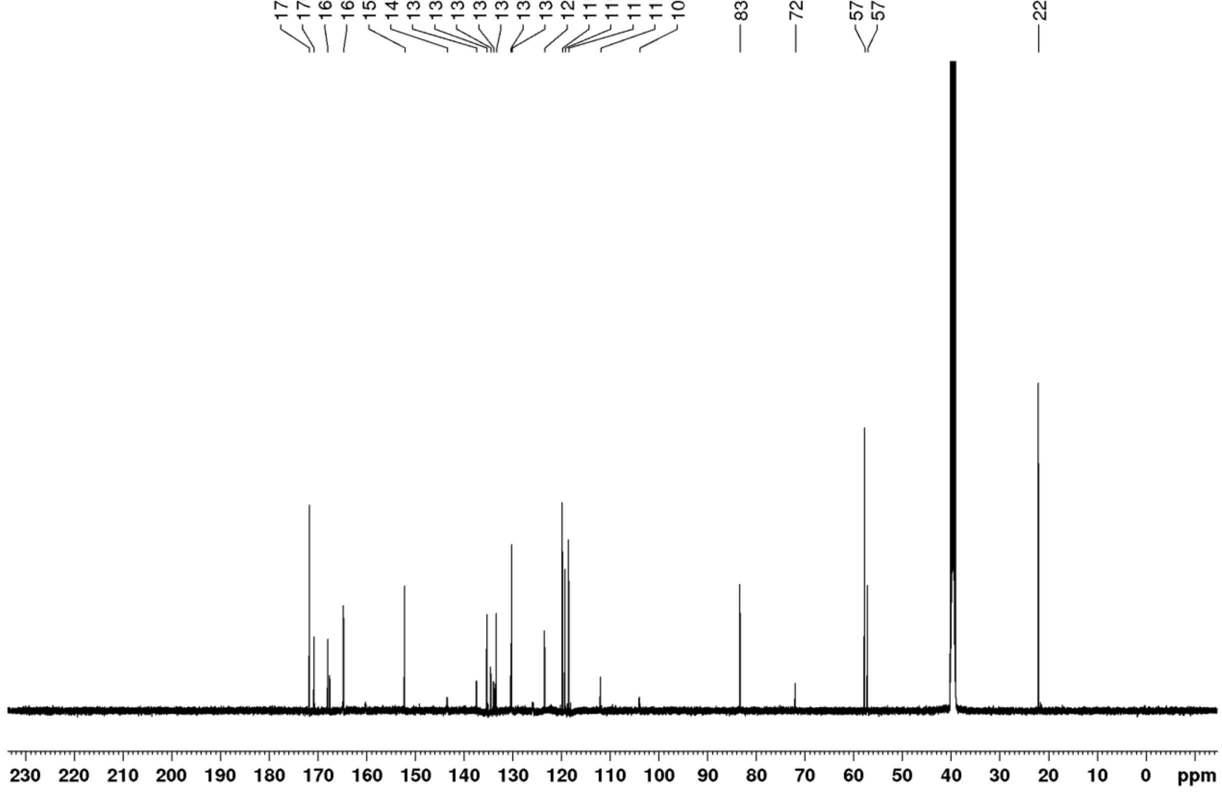
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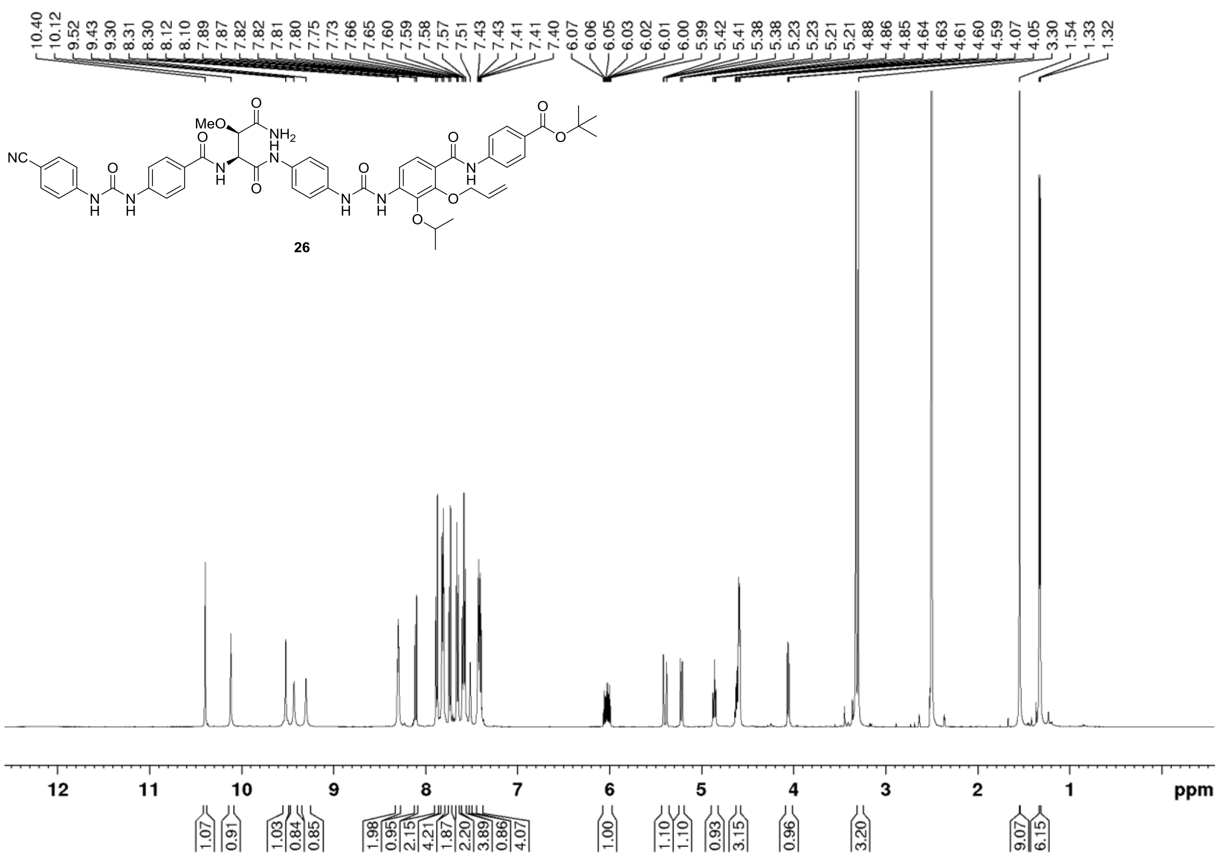
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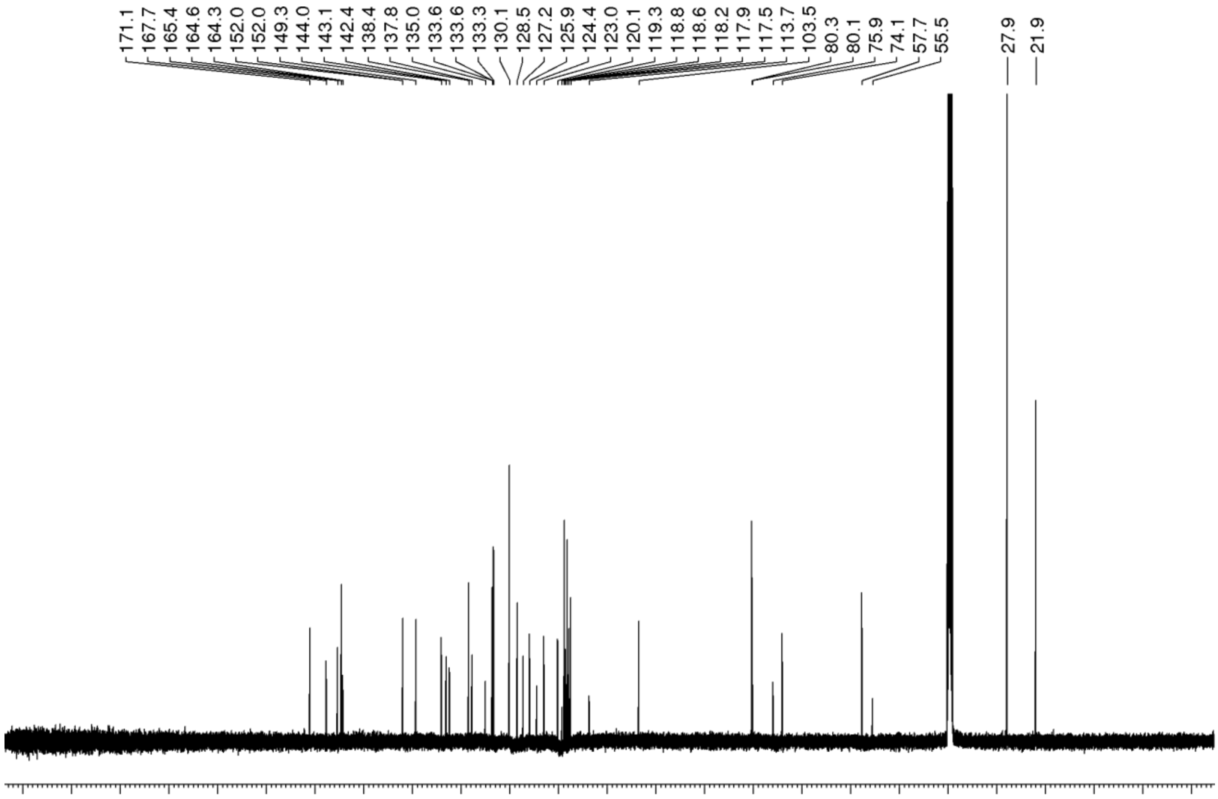
26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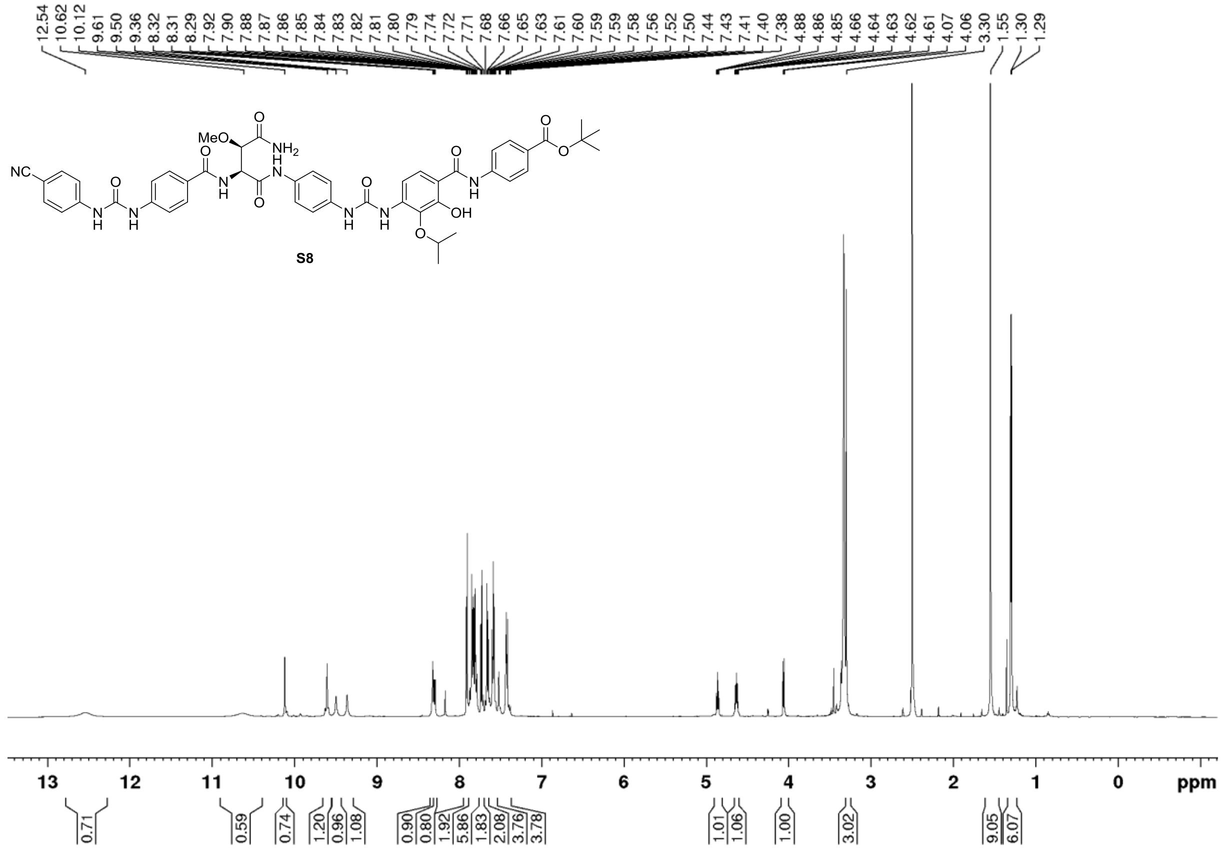
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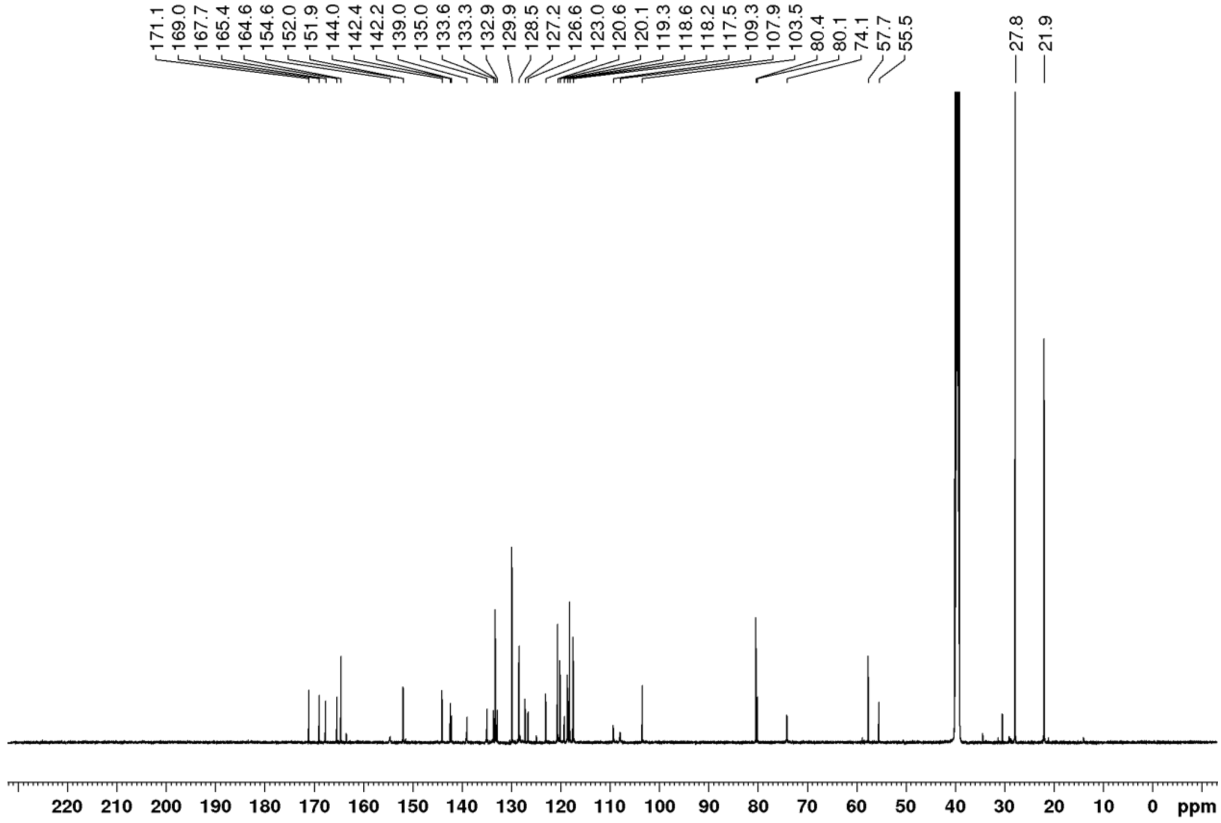
S8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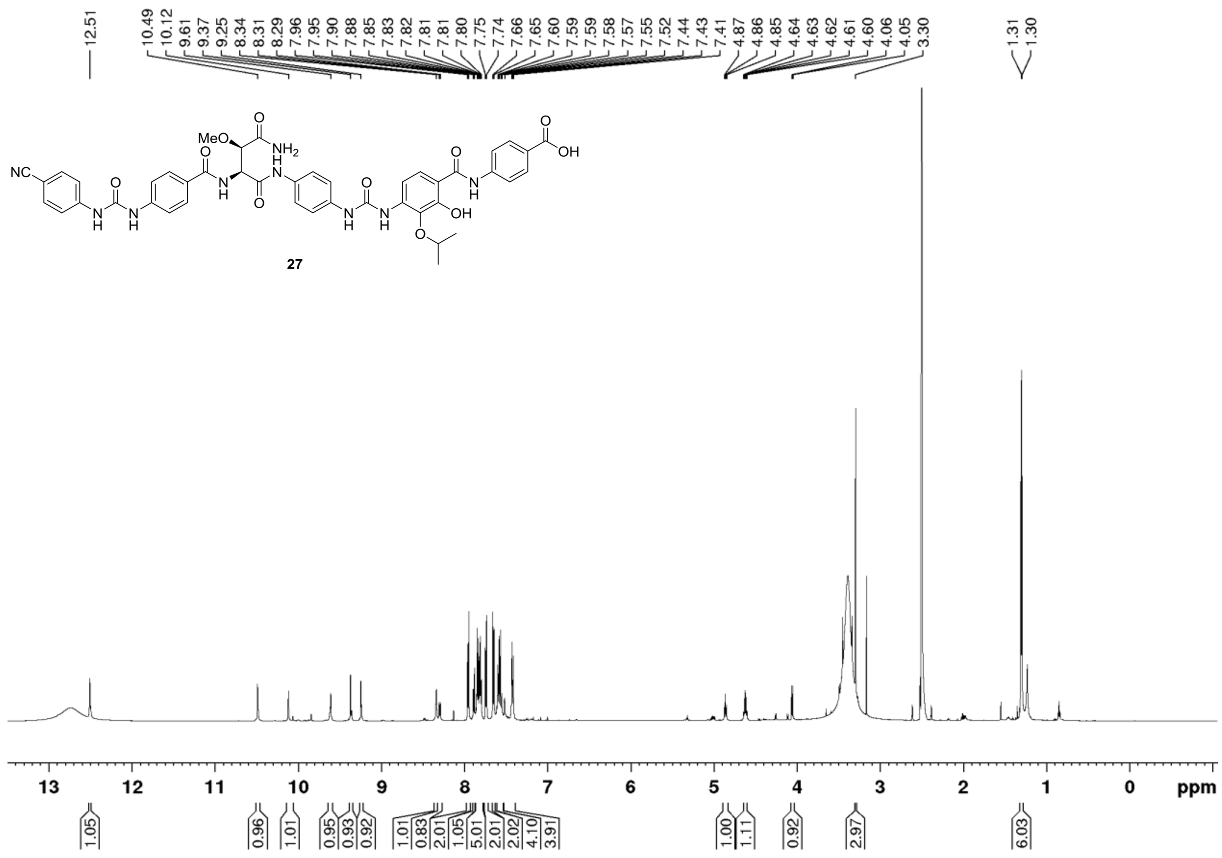
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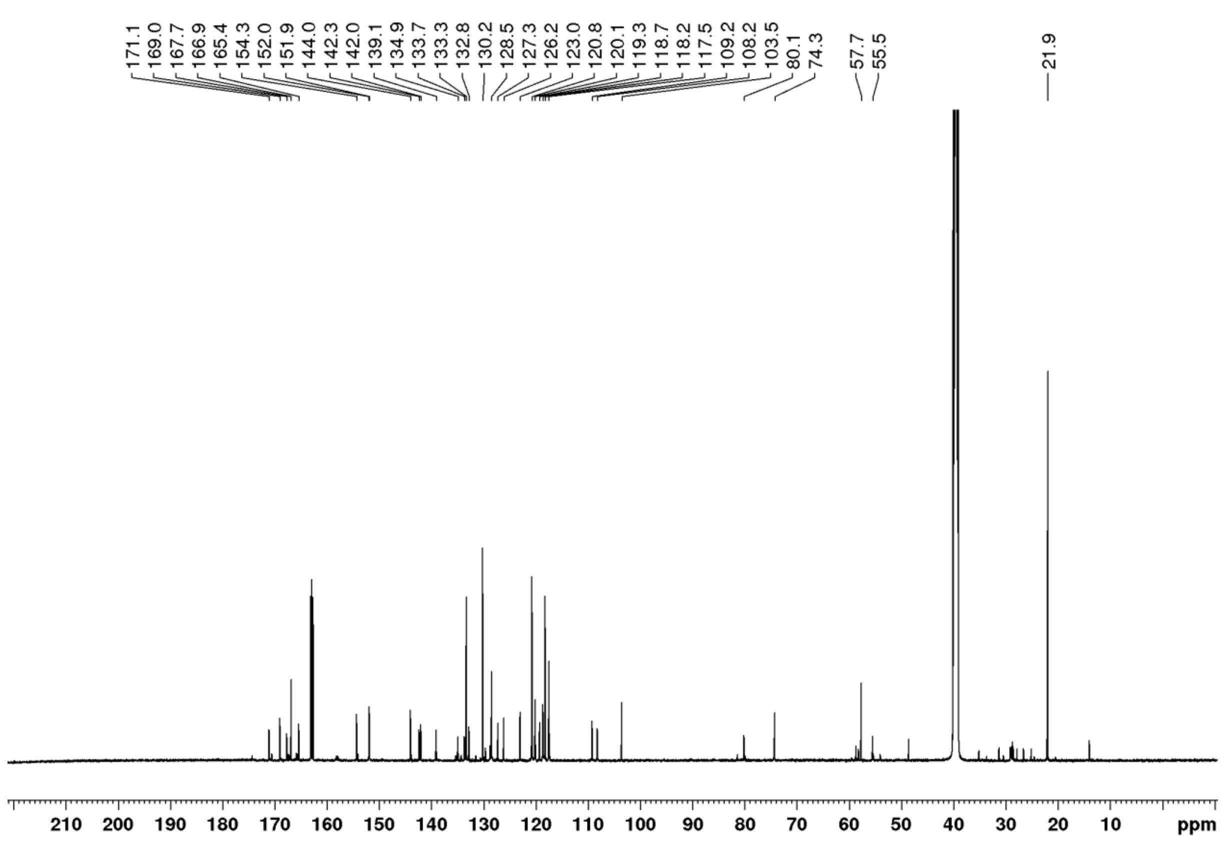
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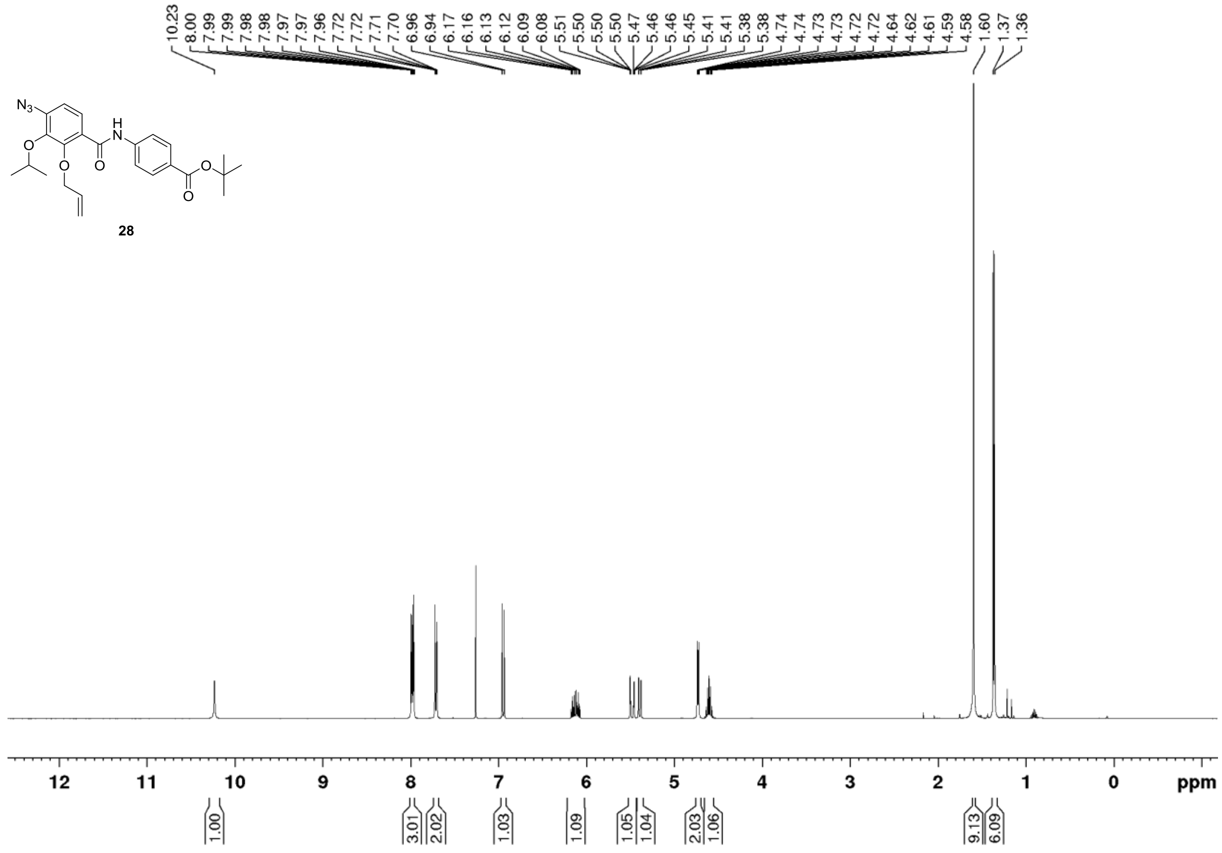

28

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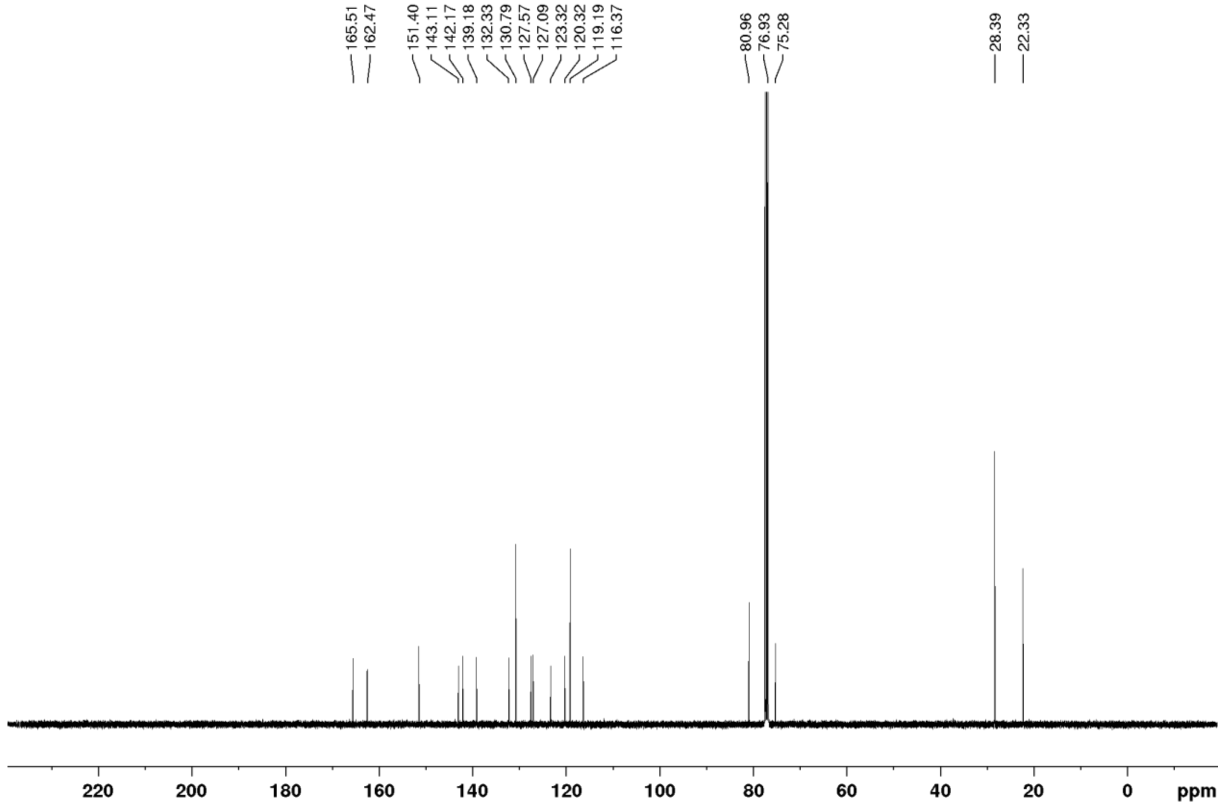

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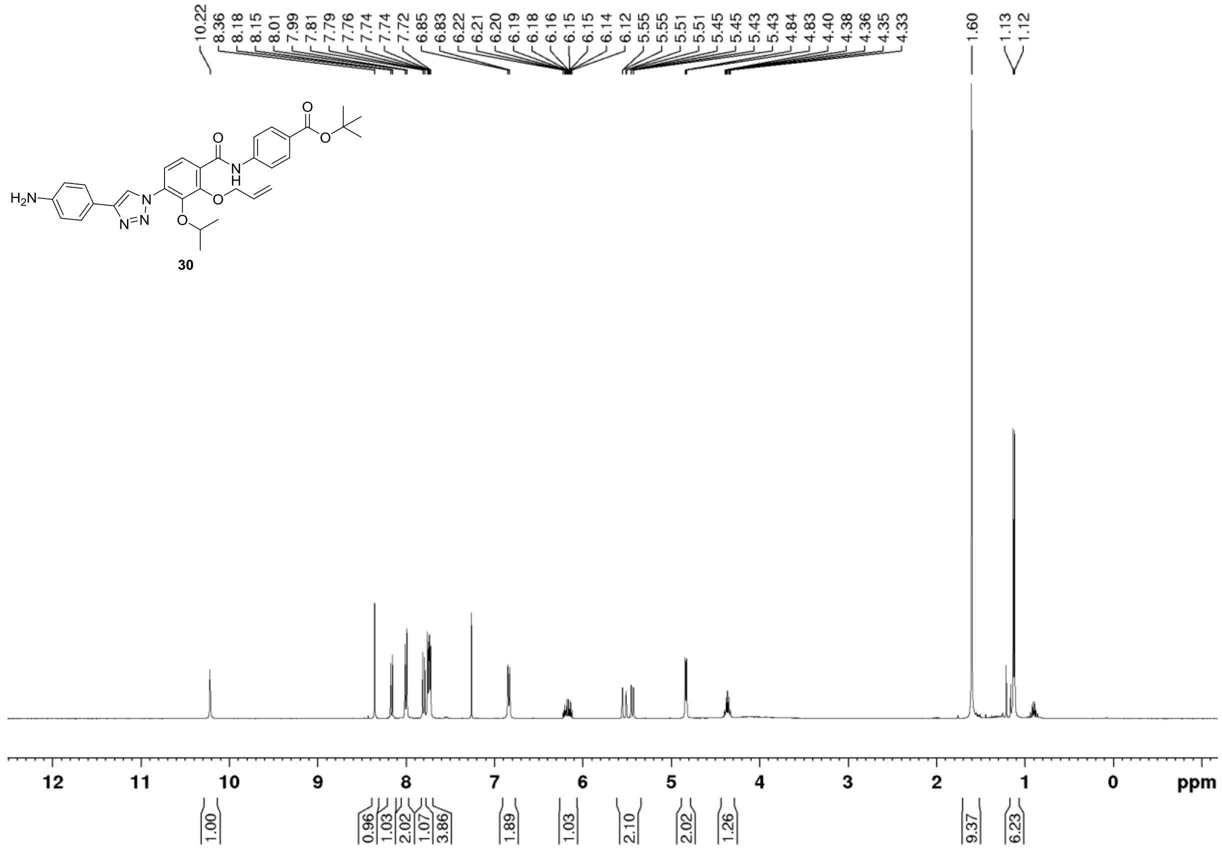
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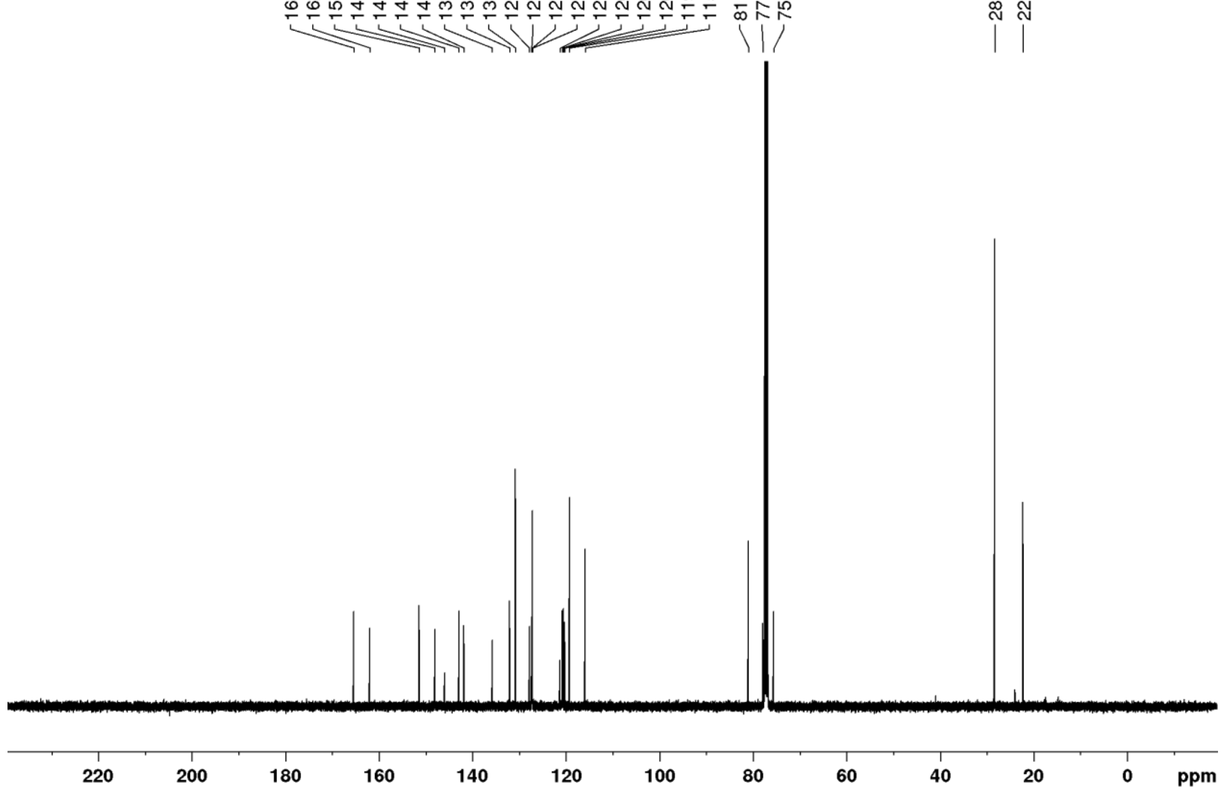
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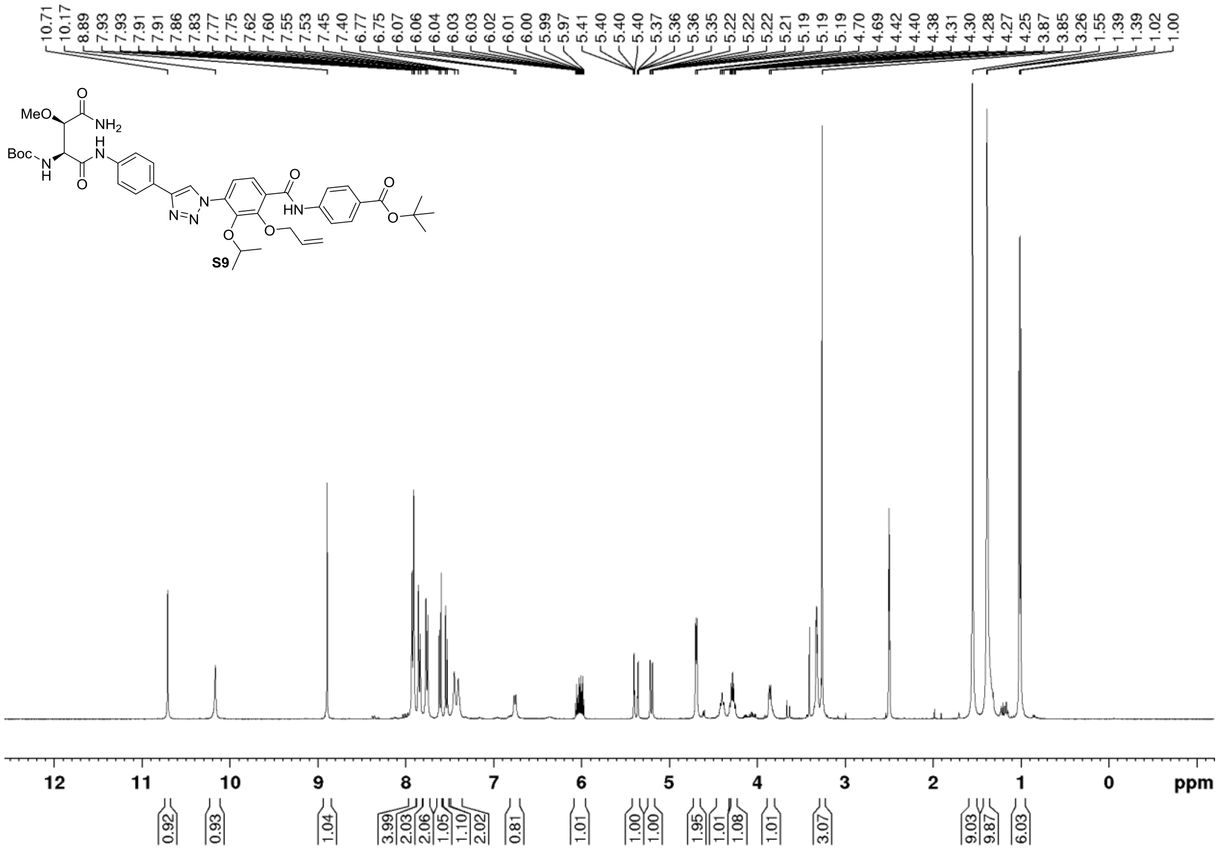
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6.03
6.03
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6.00
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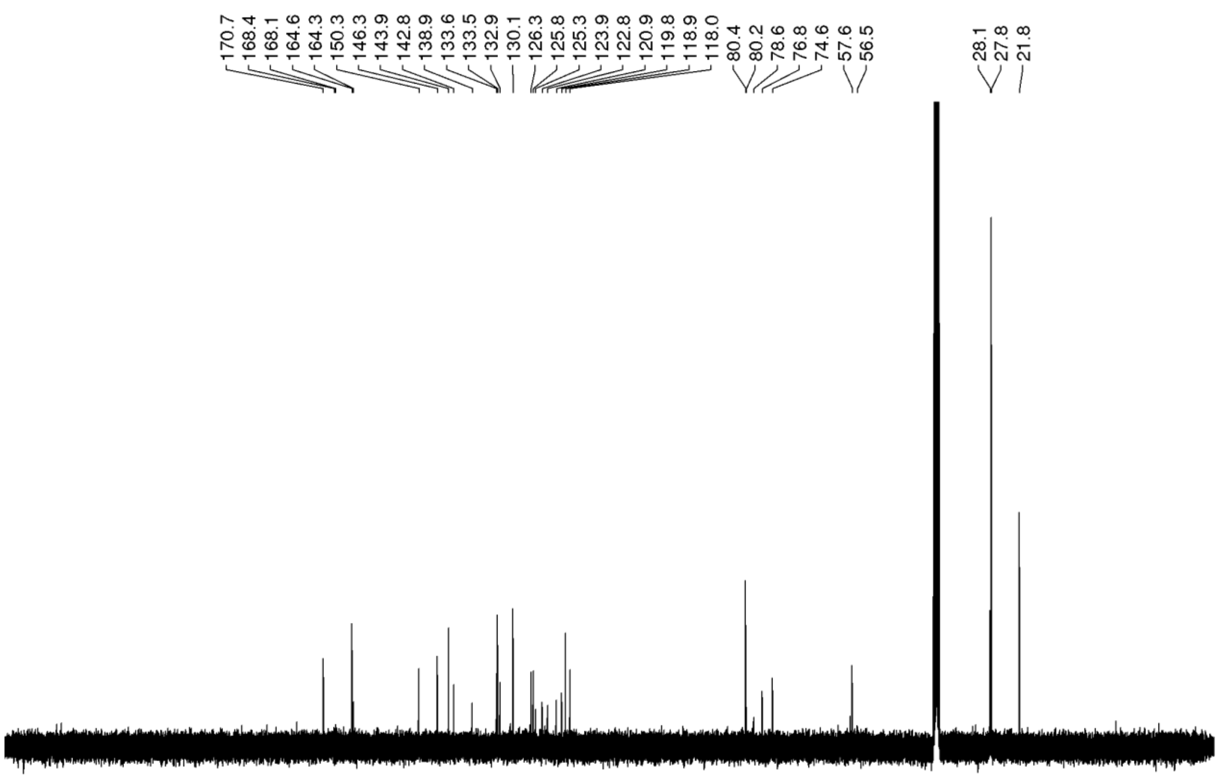
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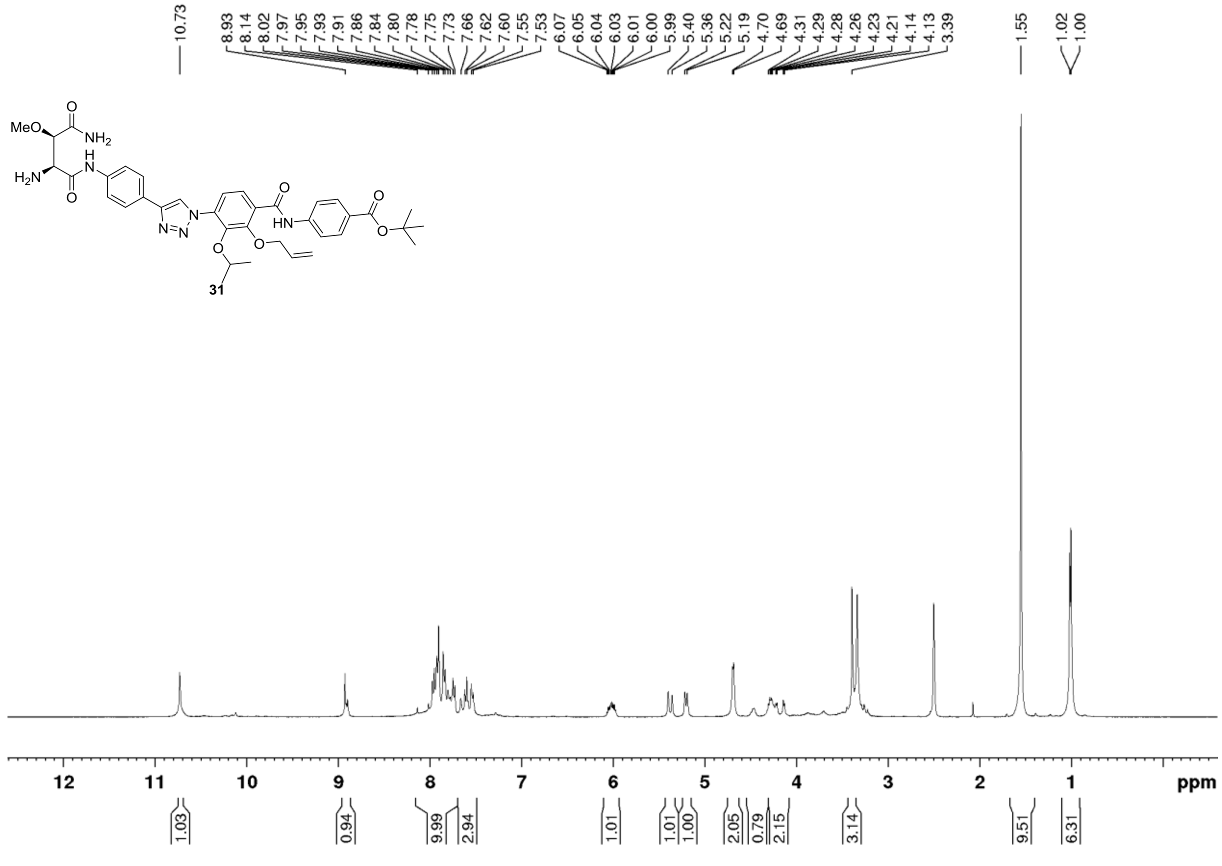
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7.91
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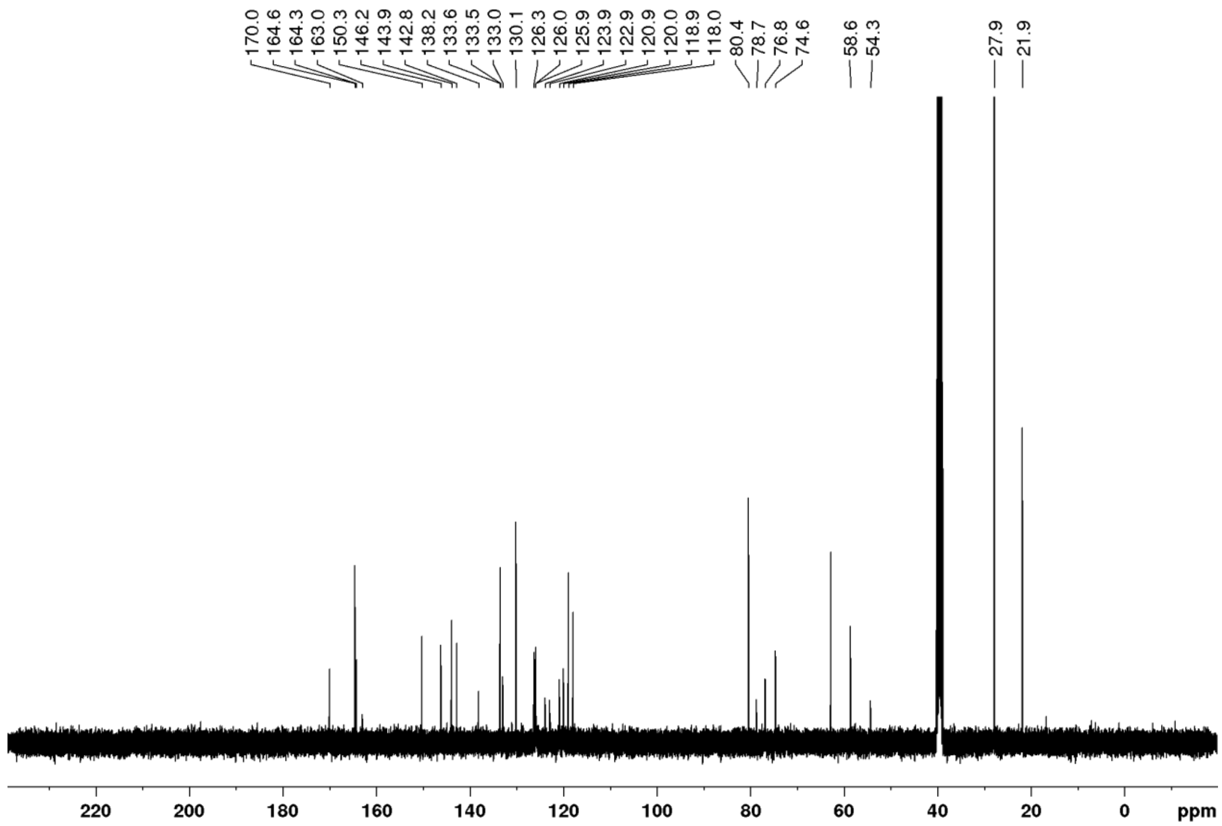
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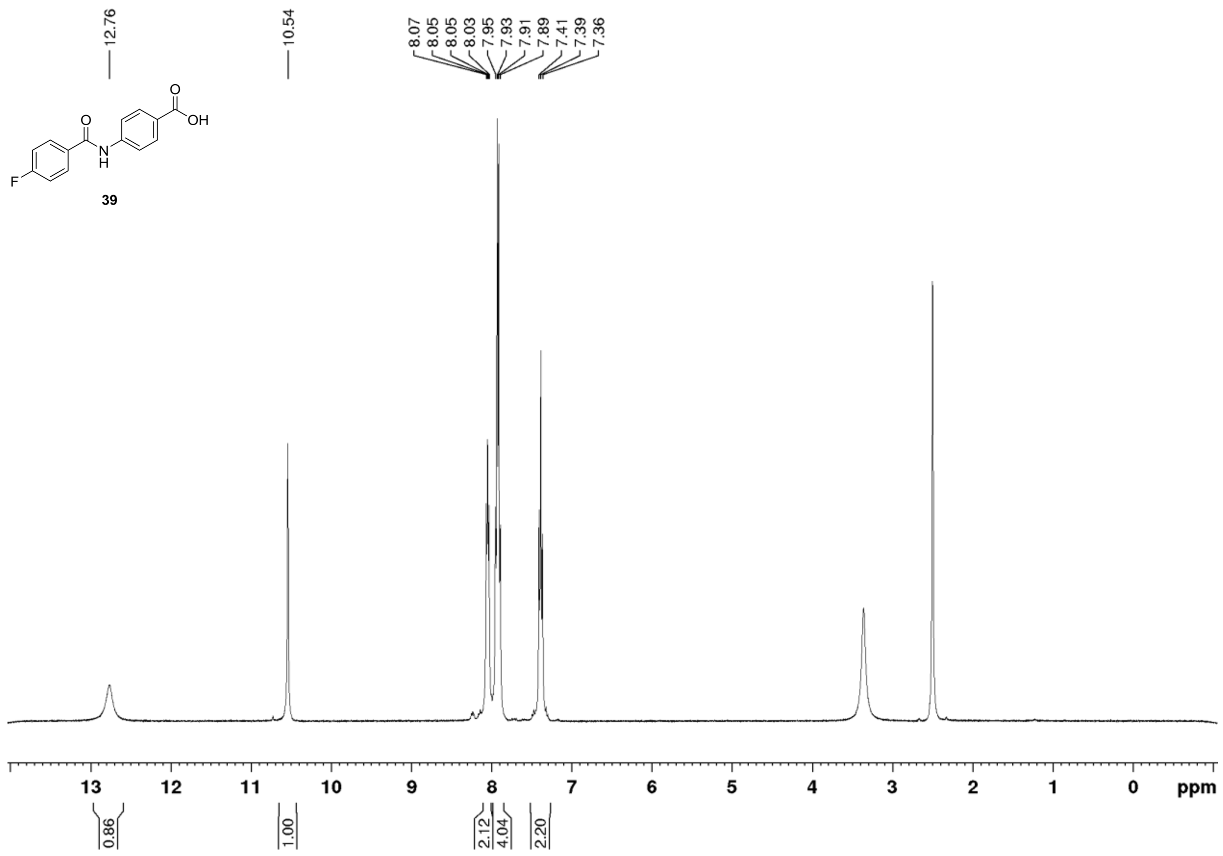
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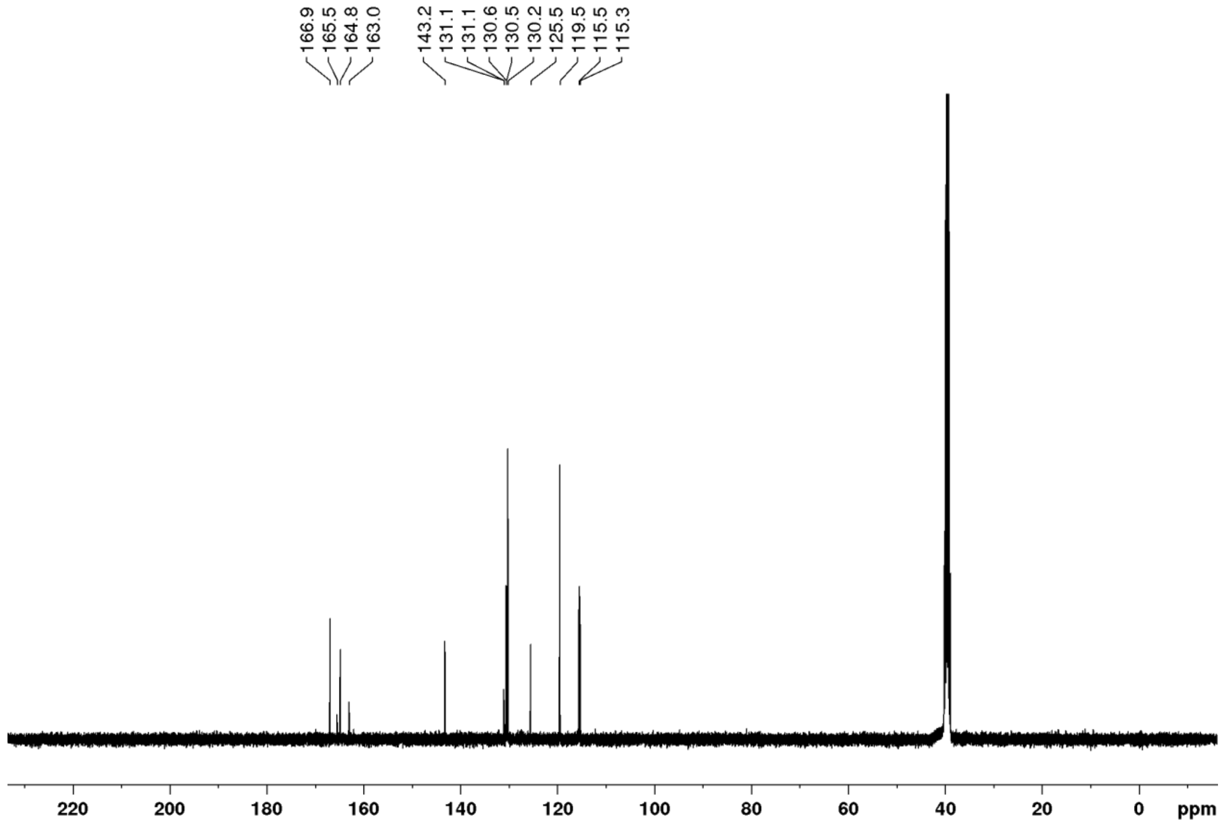
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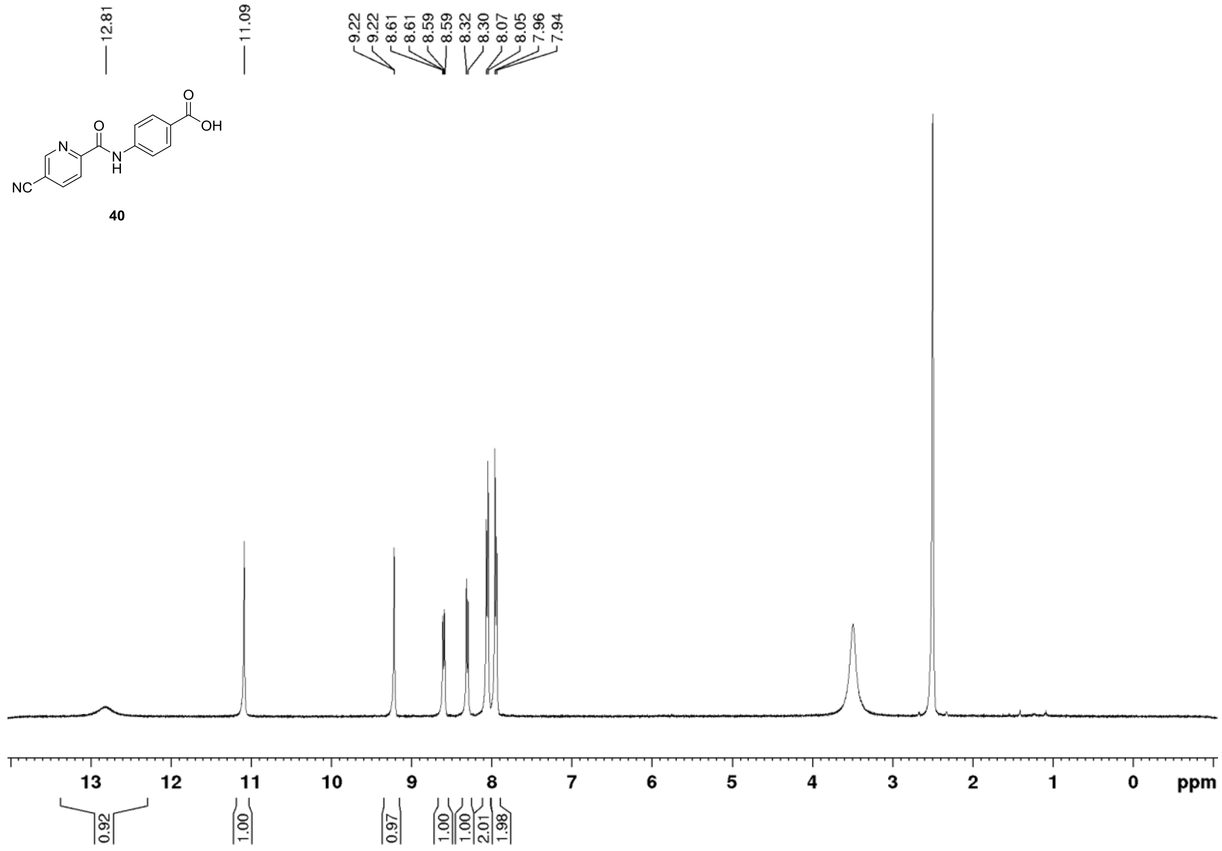

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8.59
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7.94
13
12
11
10
9
8
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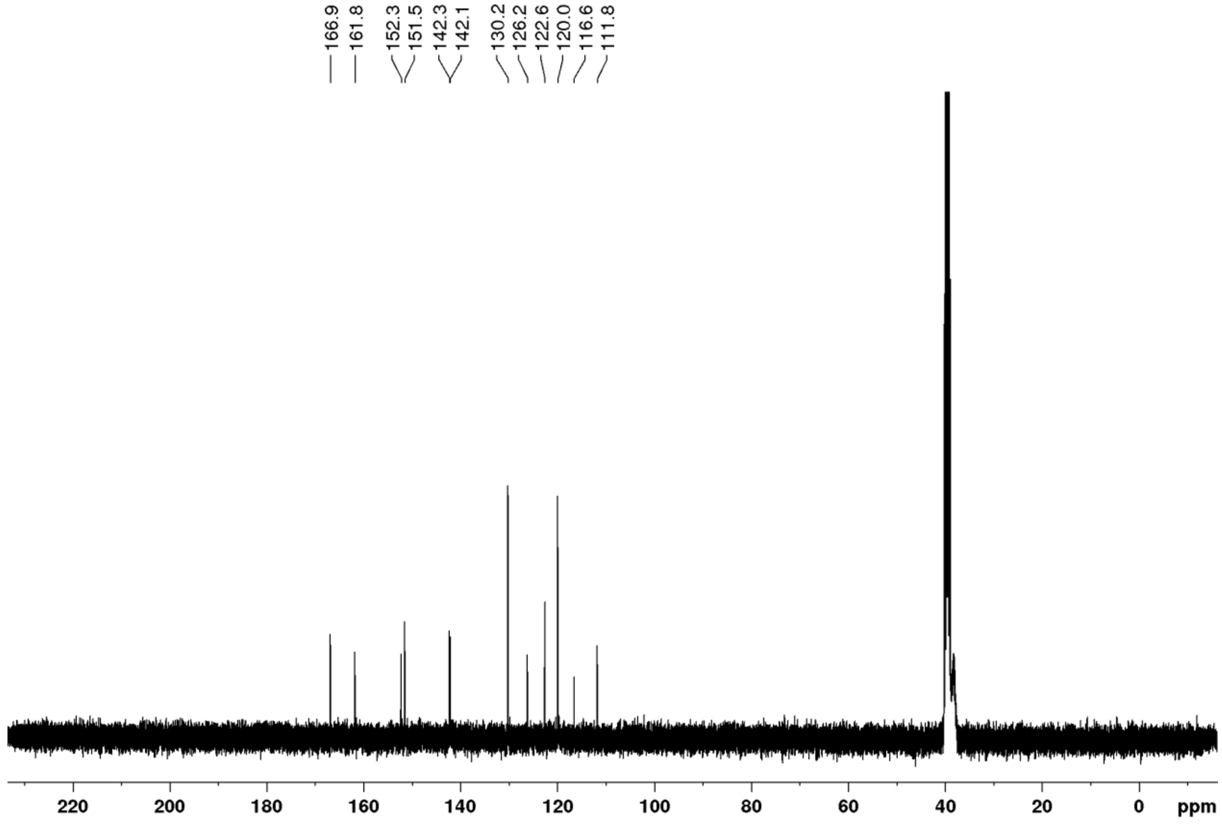

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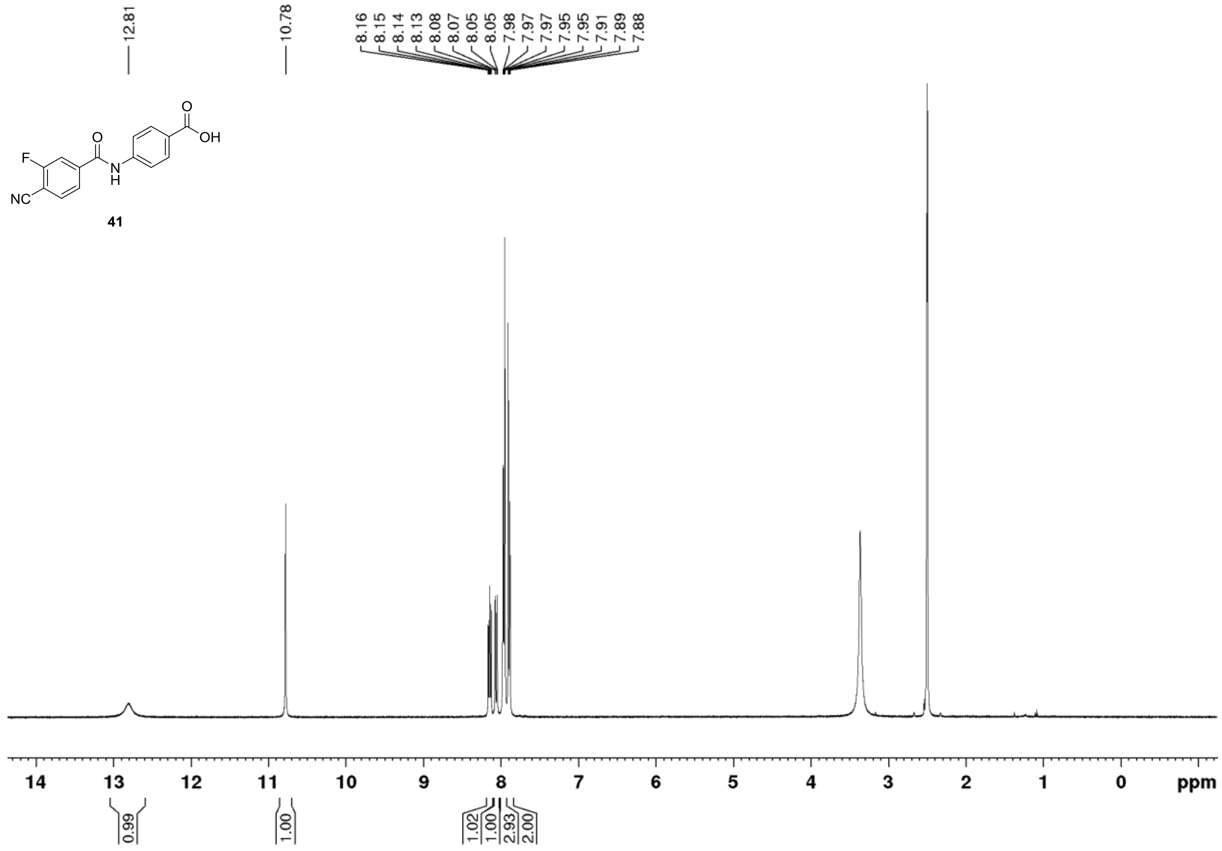

41

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O
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H
O
OH
NC
14
13
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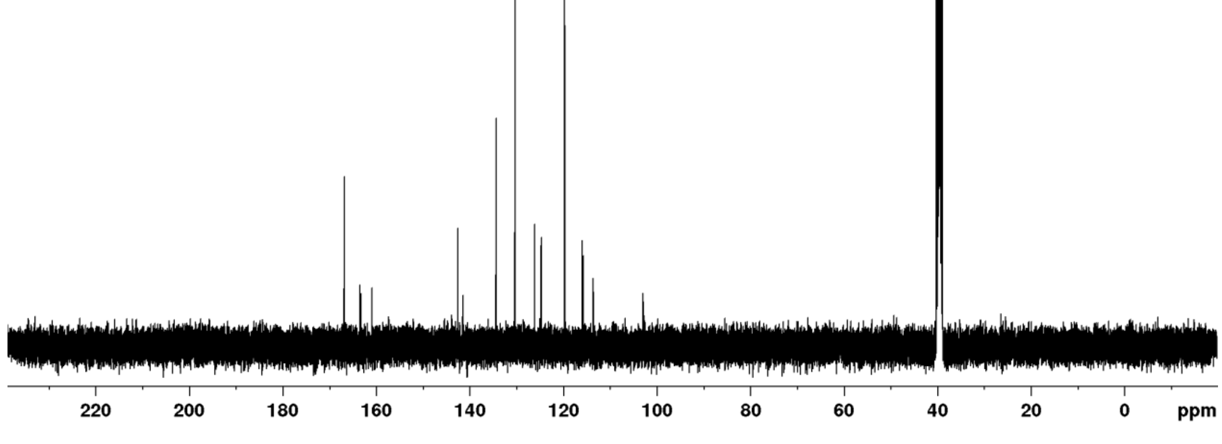

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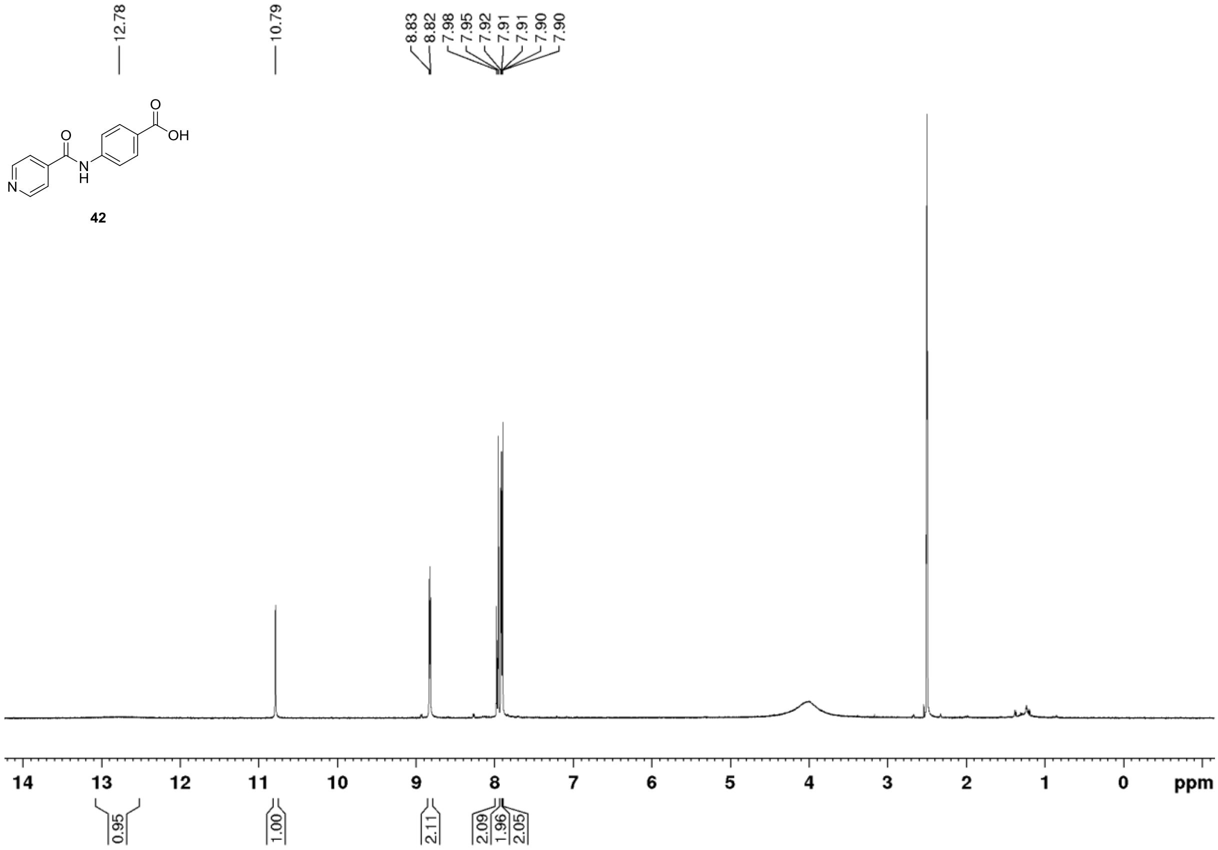

42

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H
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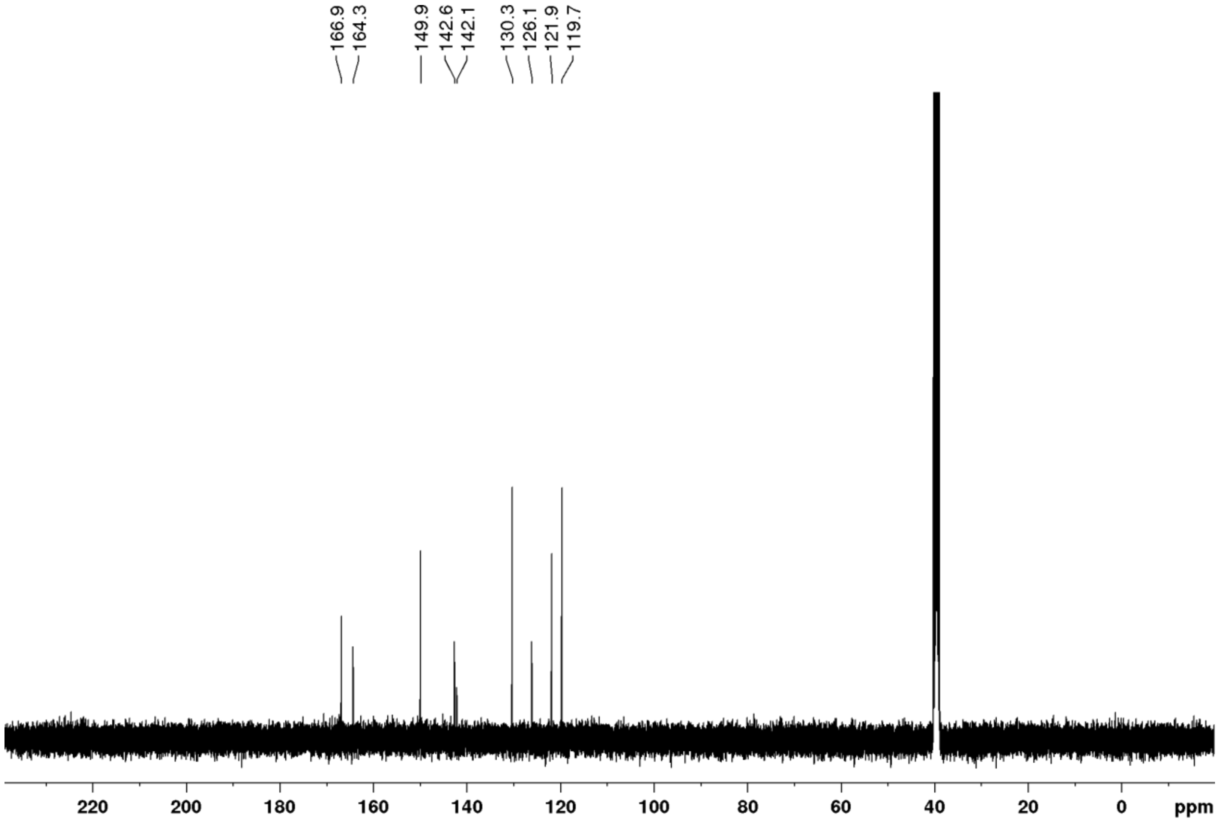

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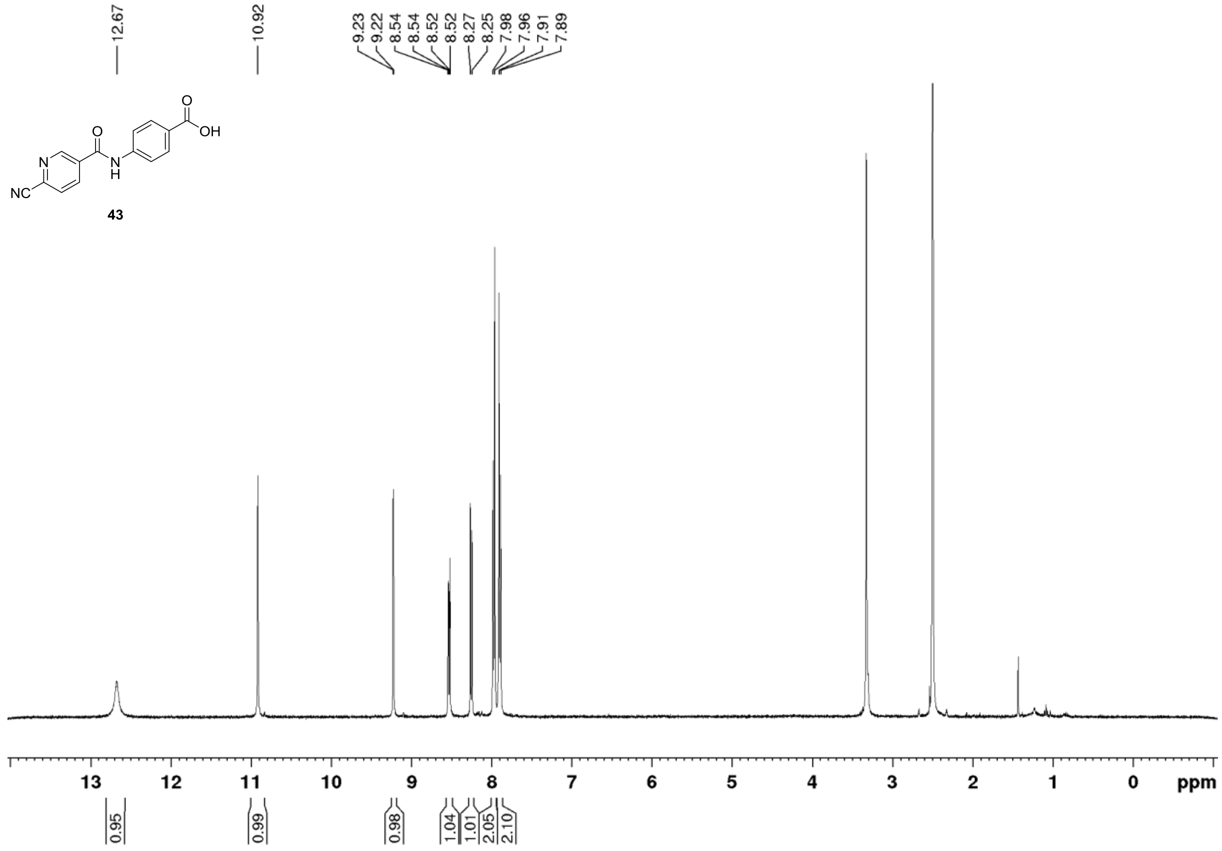
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OH
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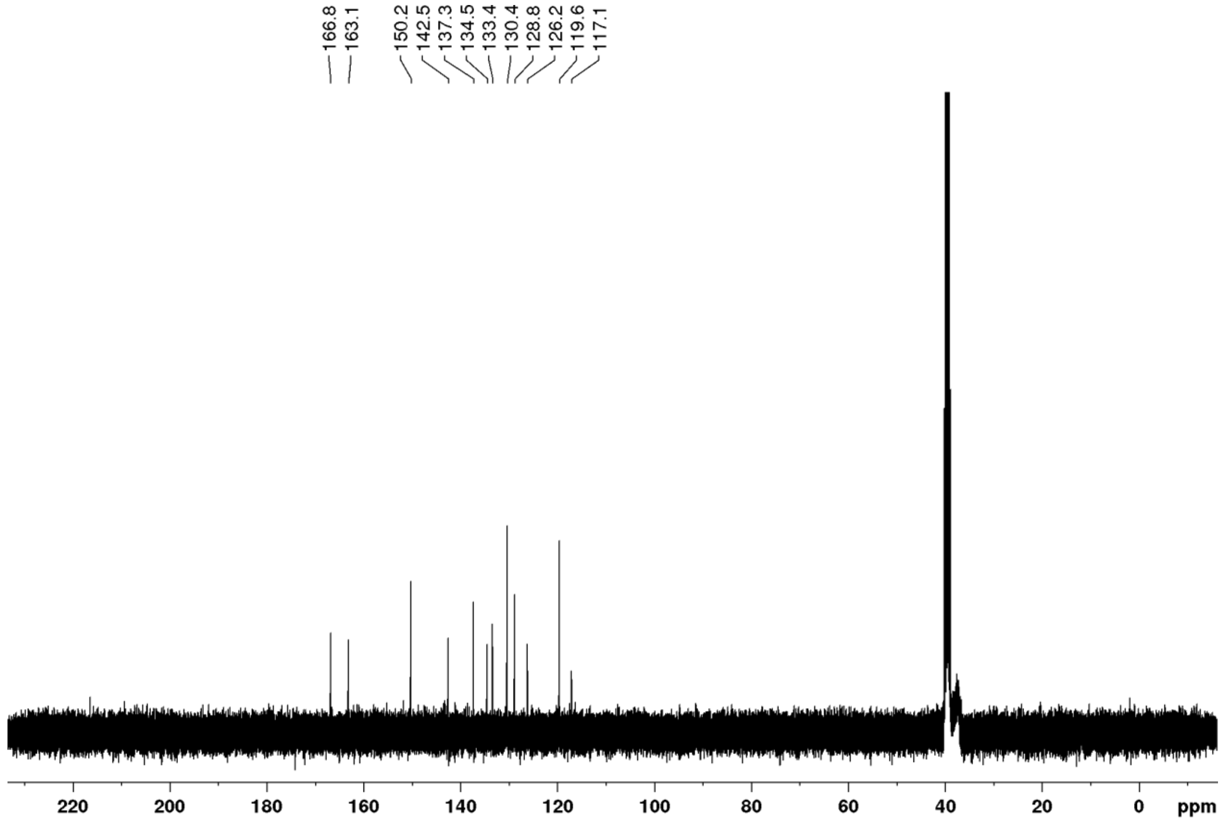
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120
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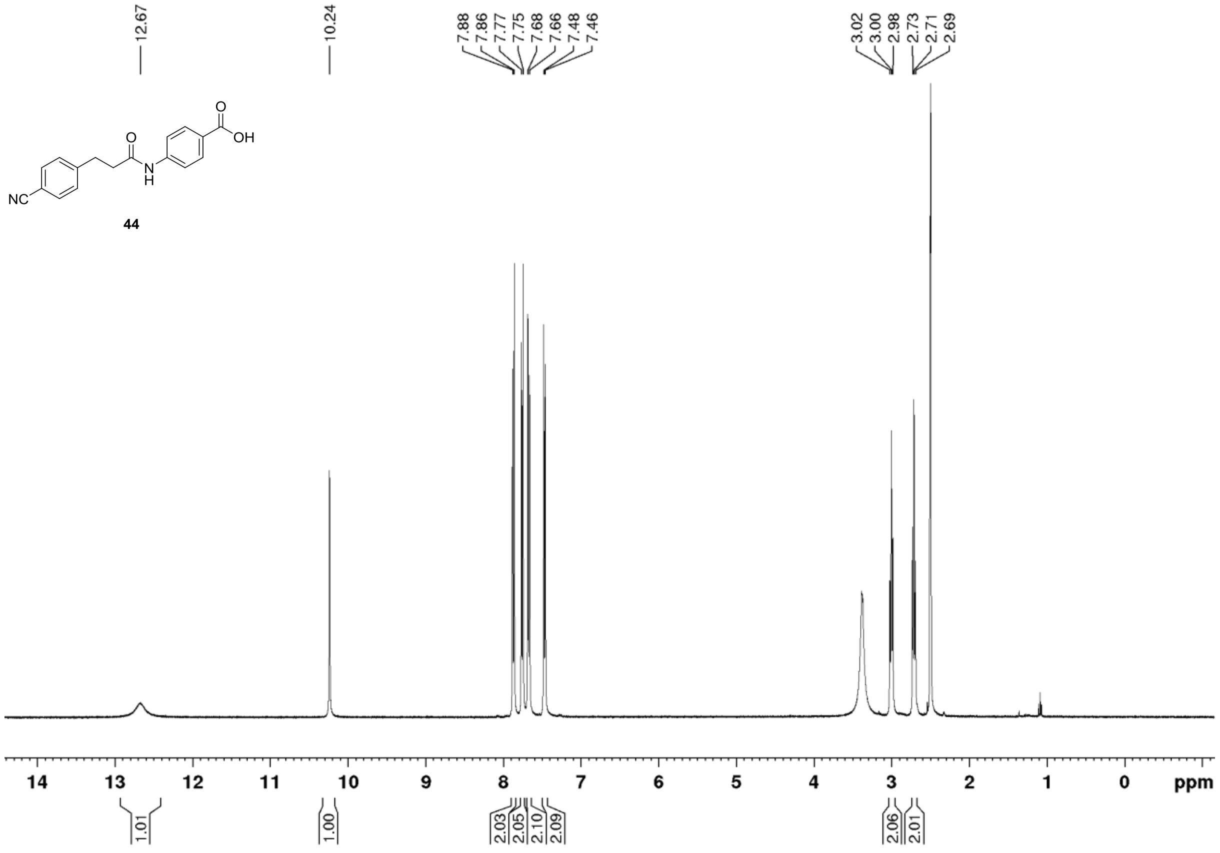
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N
H
NC
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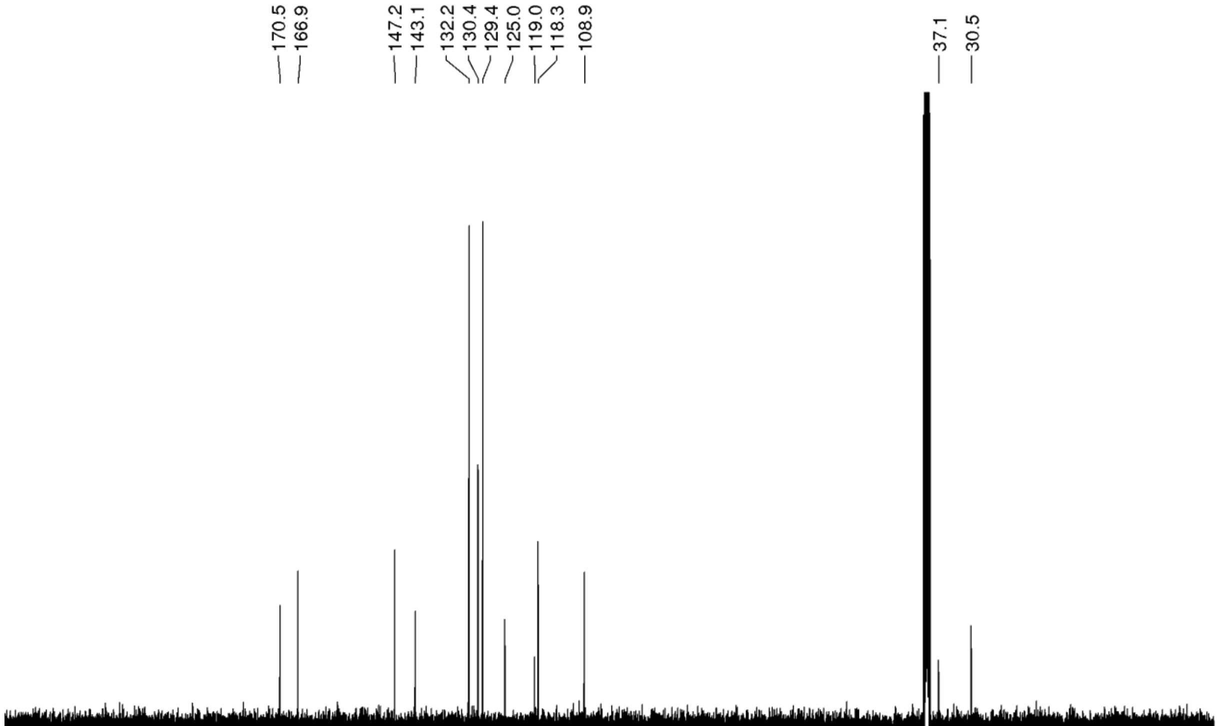
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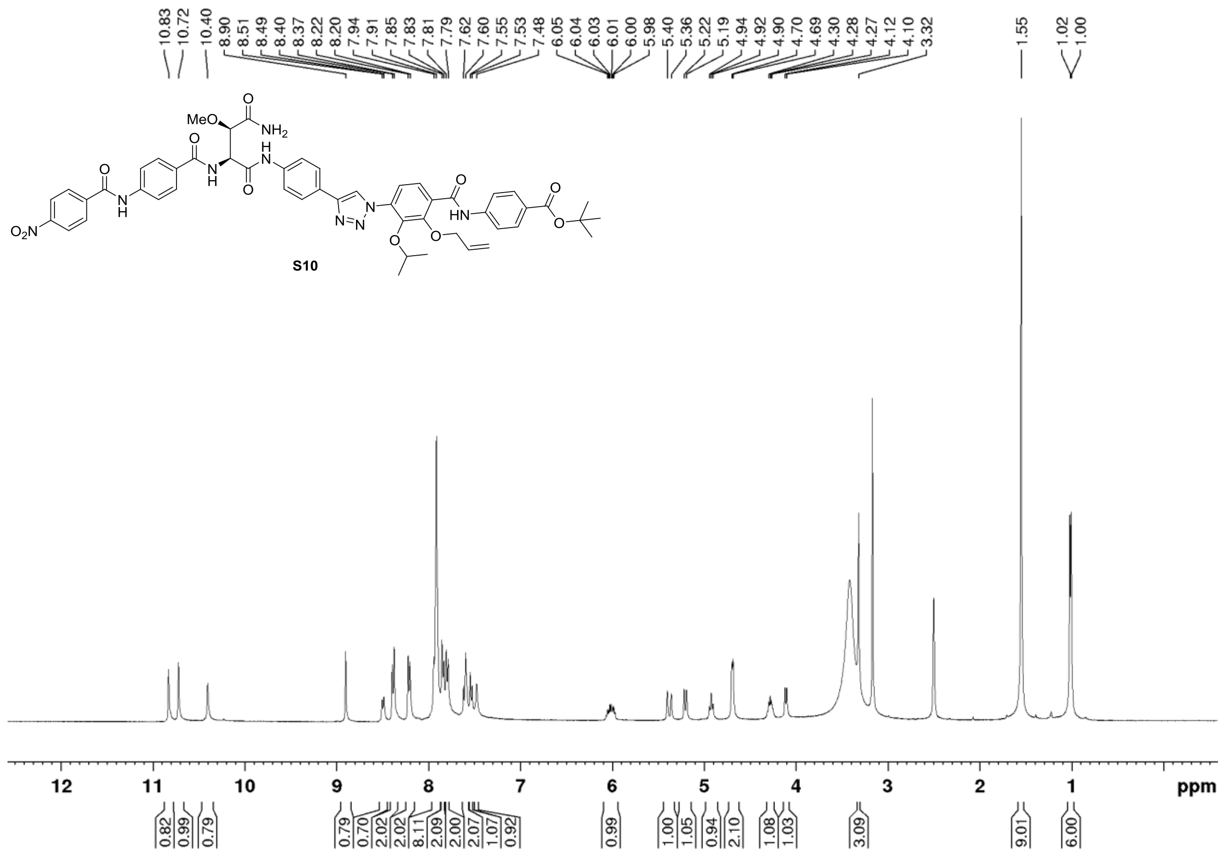
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HN
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O
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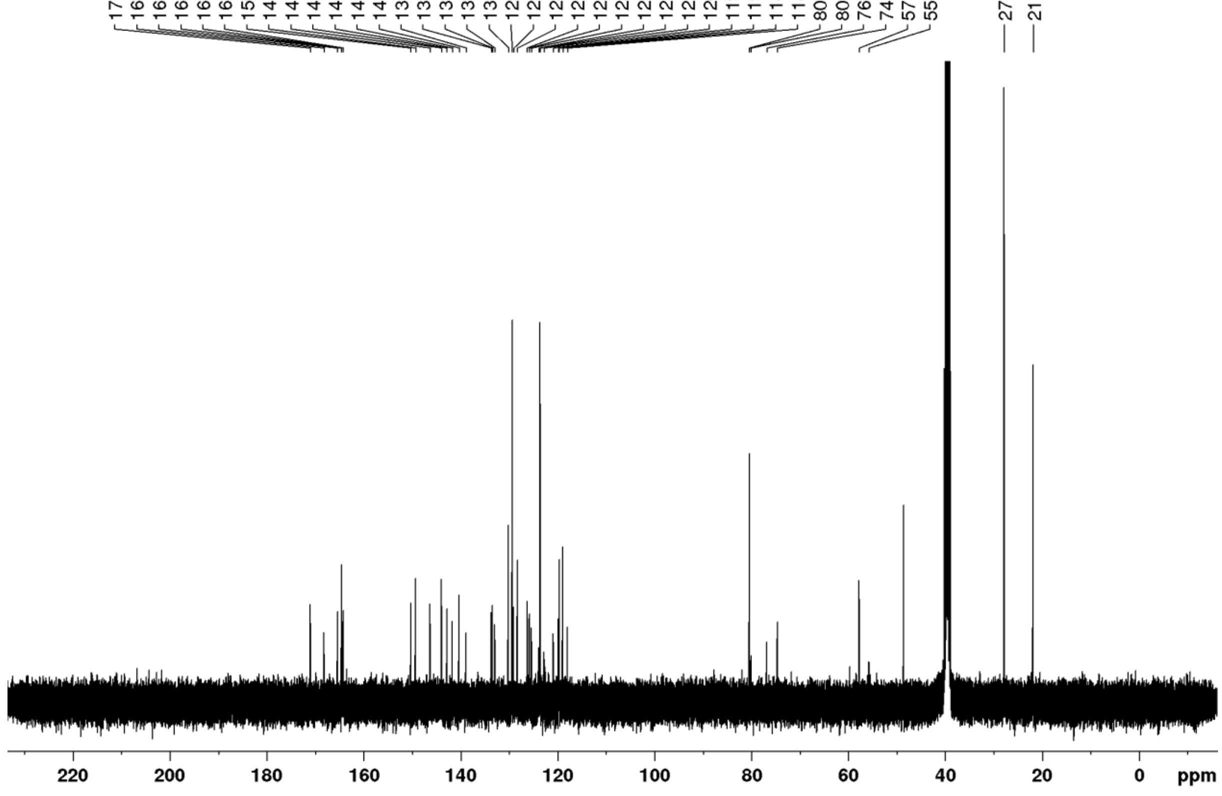
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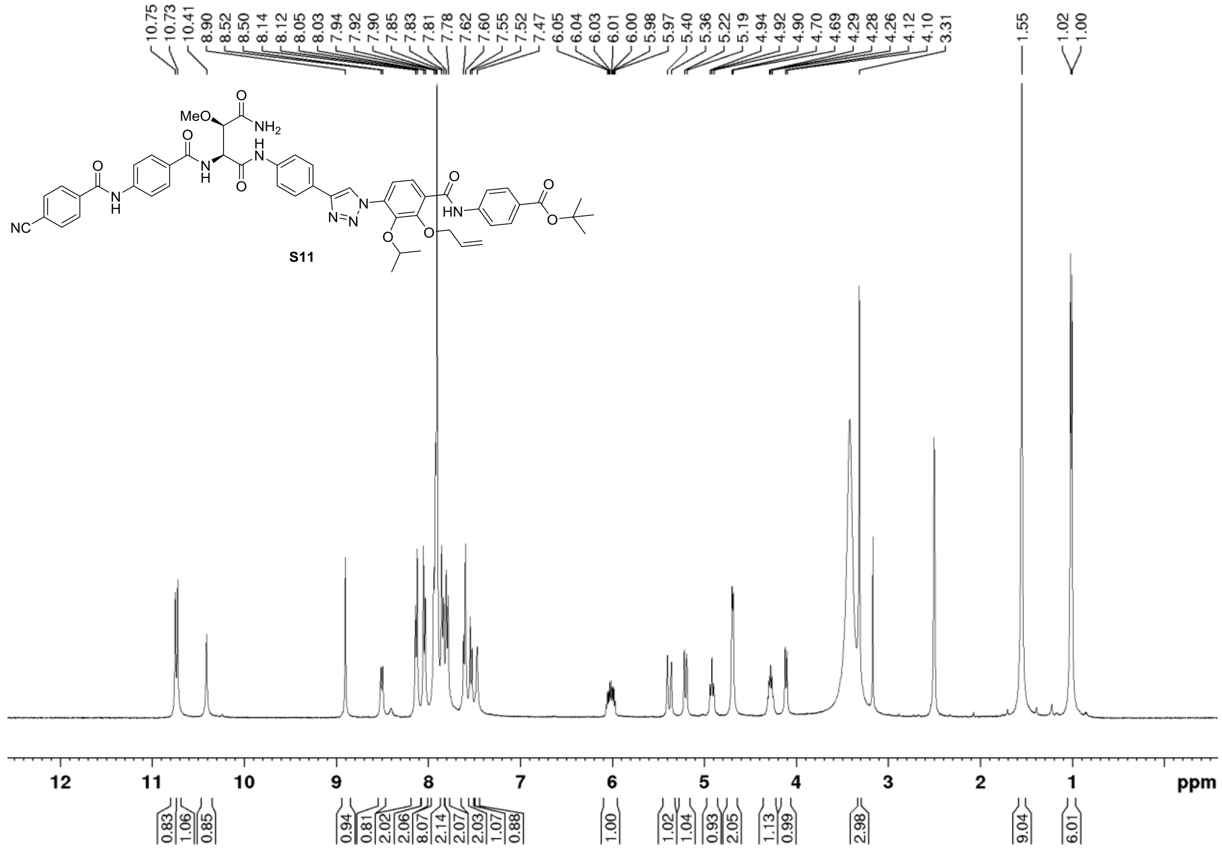
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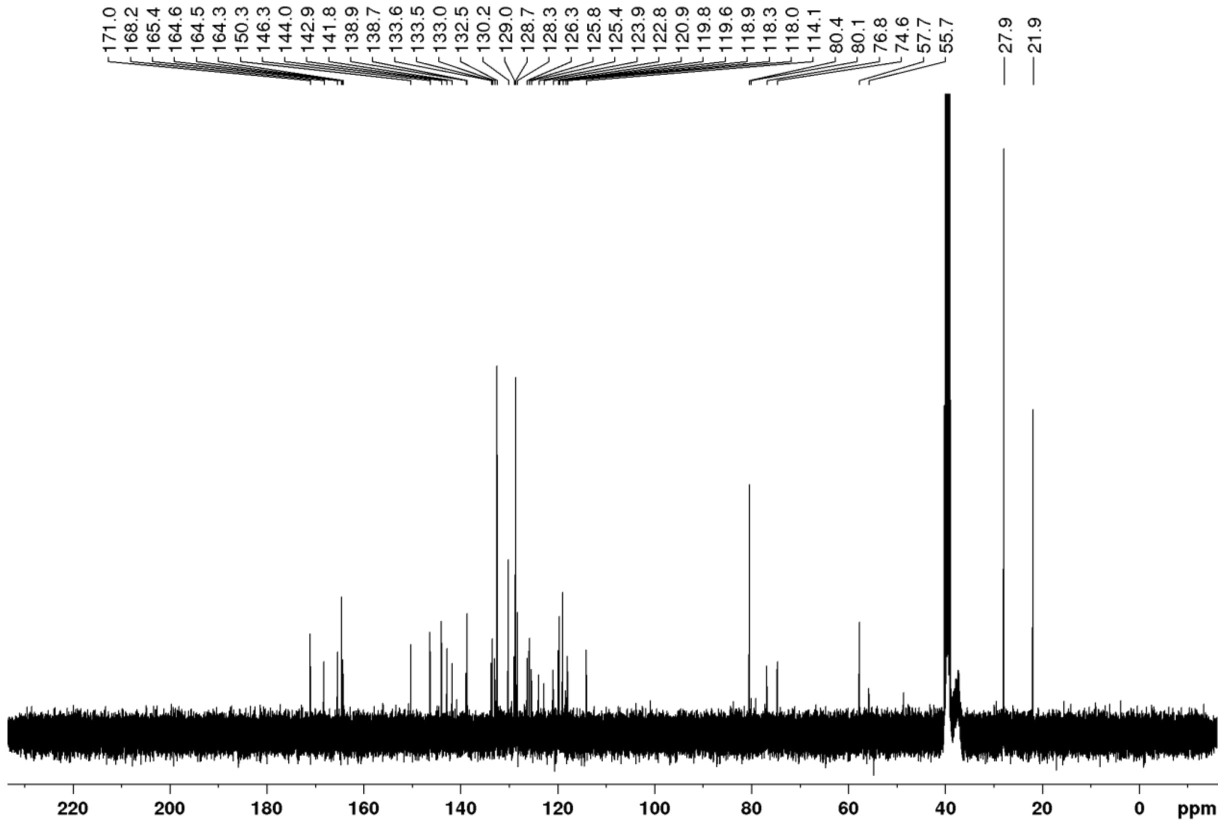
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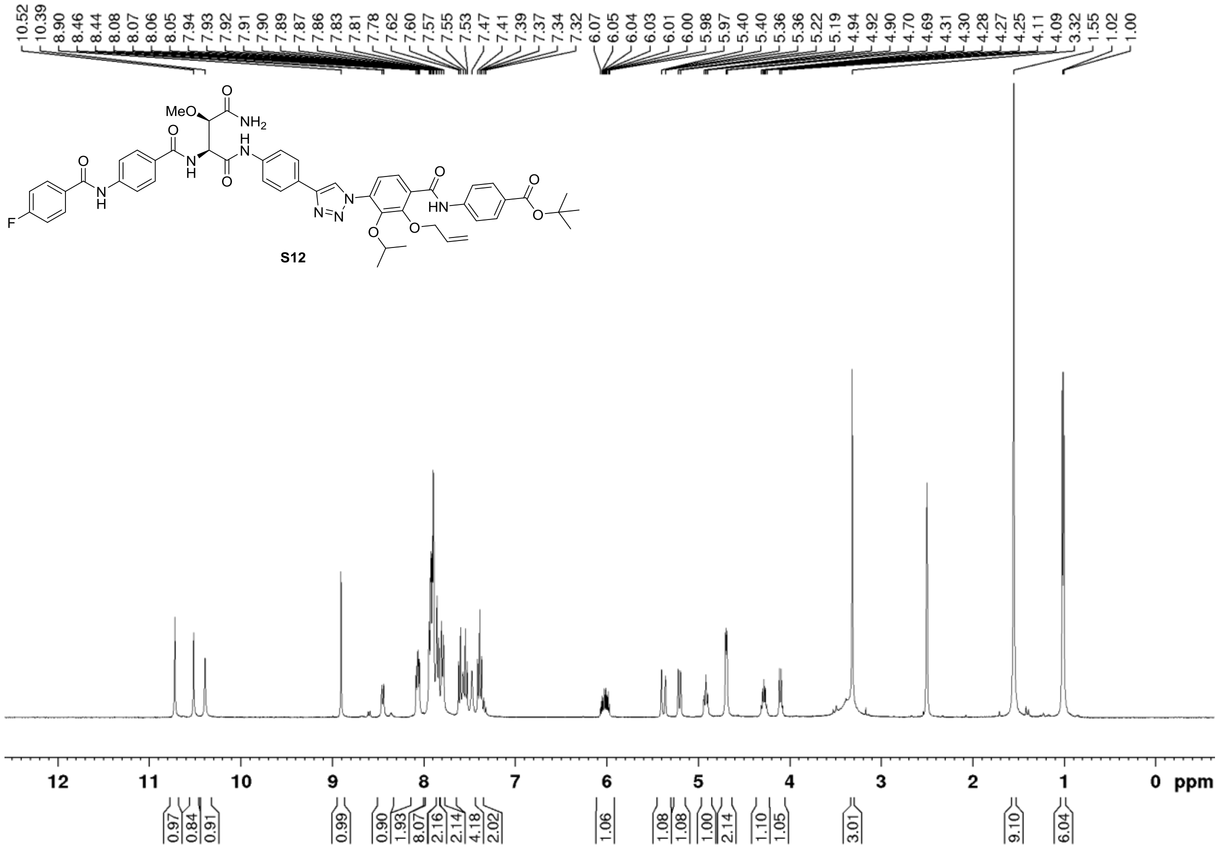
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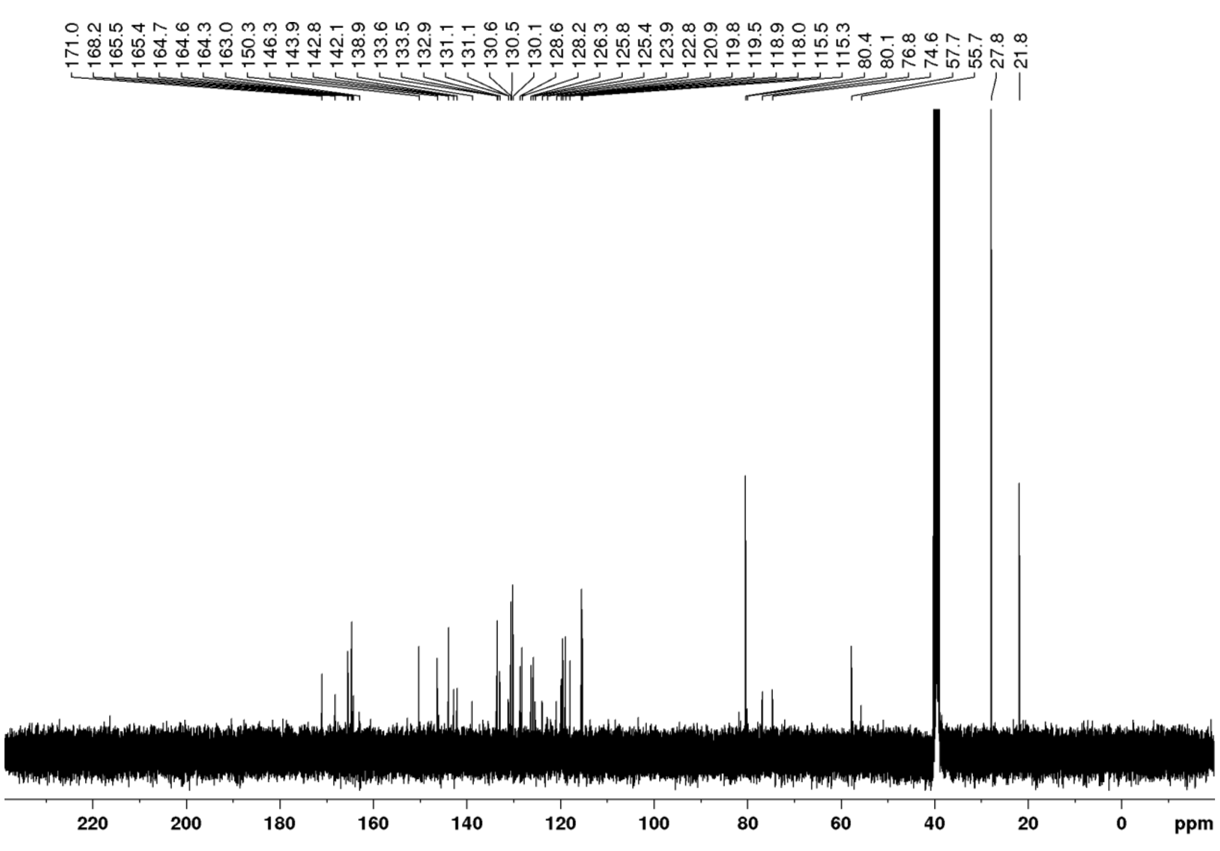
S12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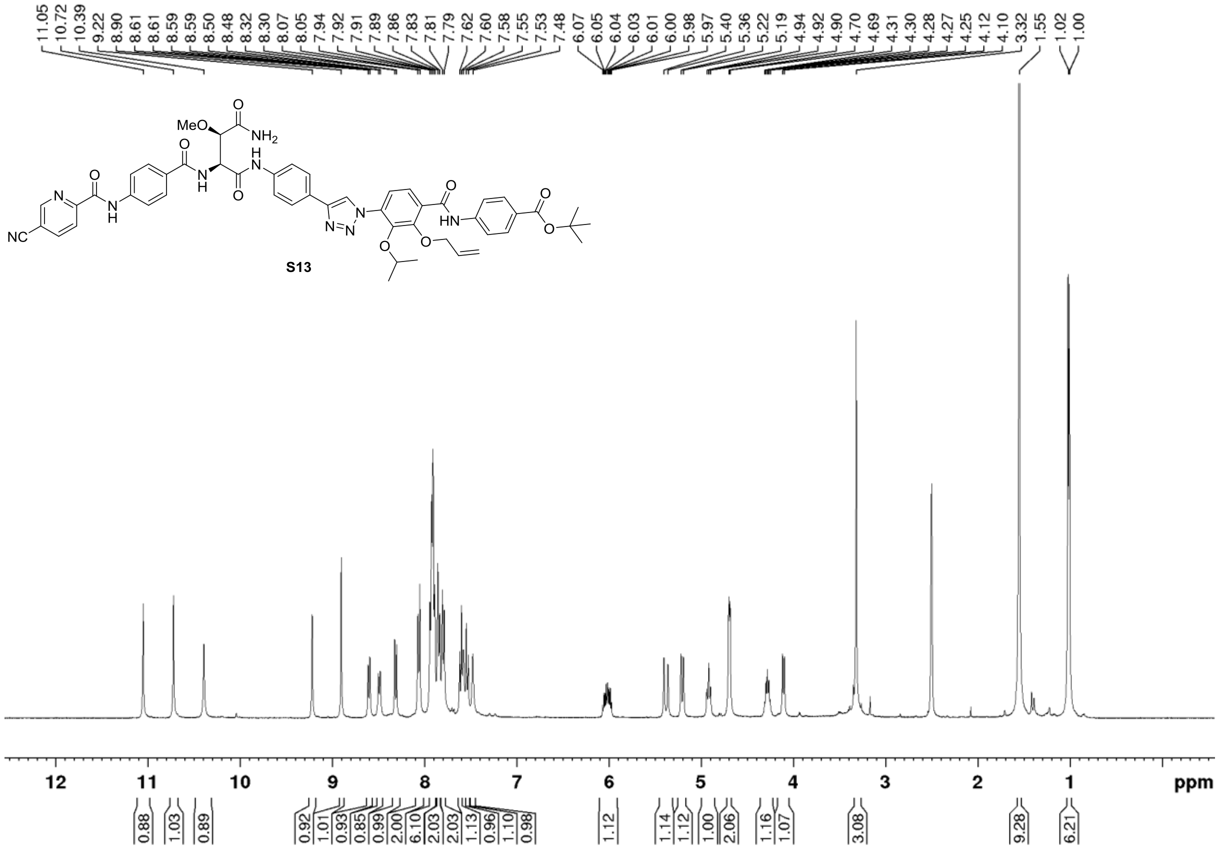
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S13
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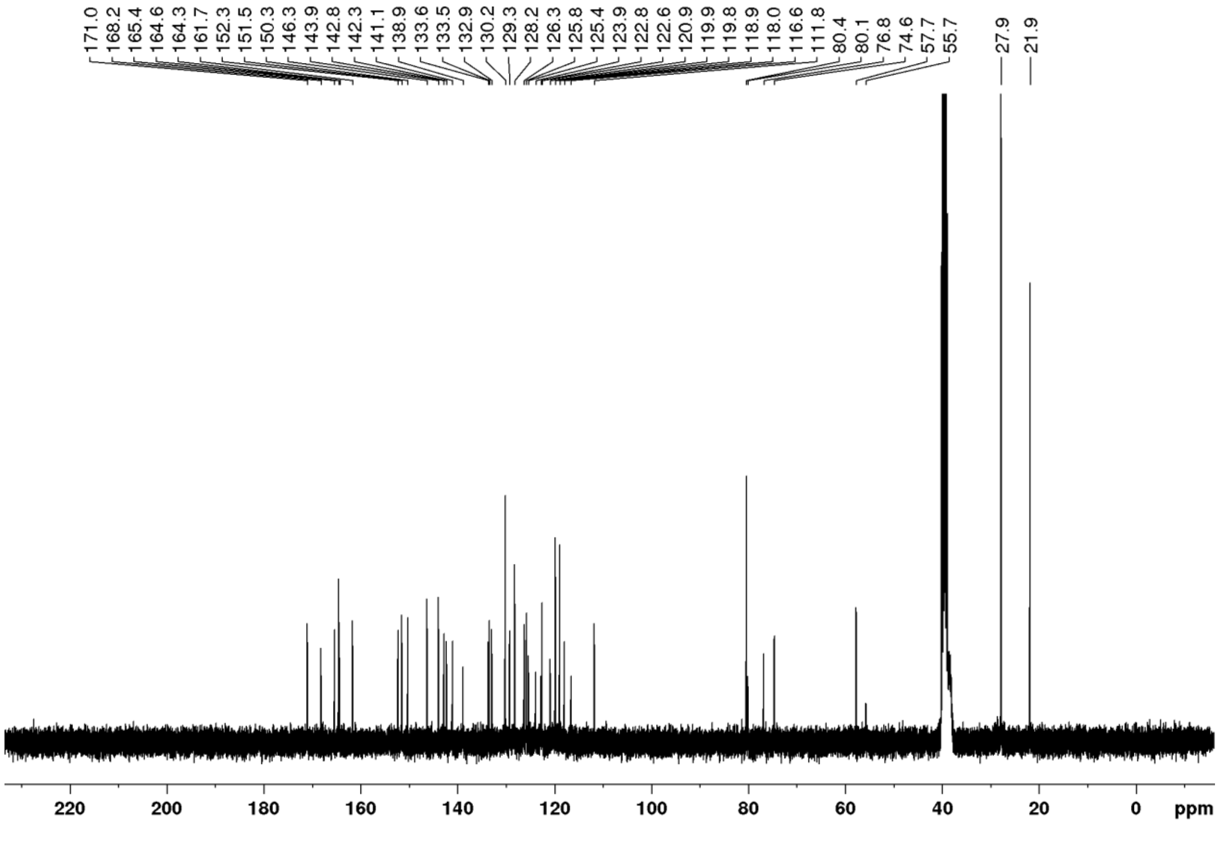
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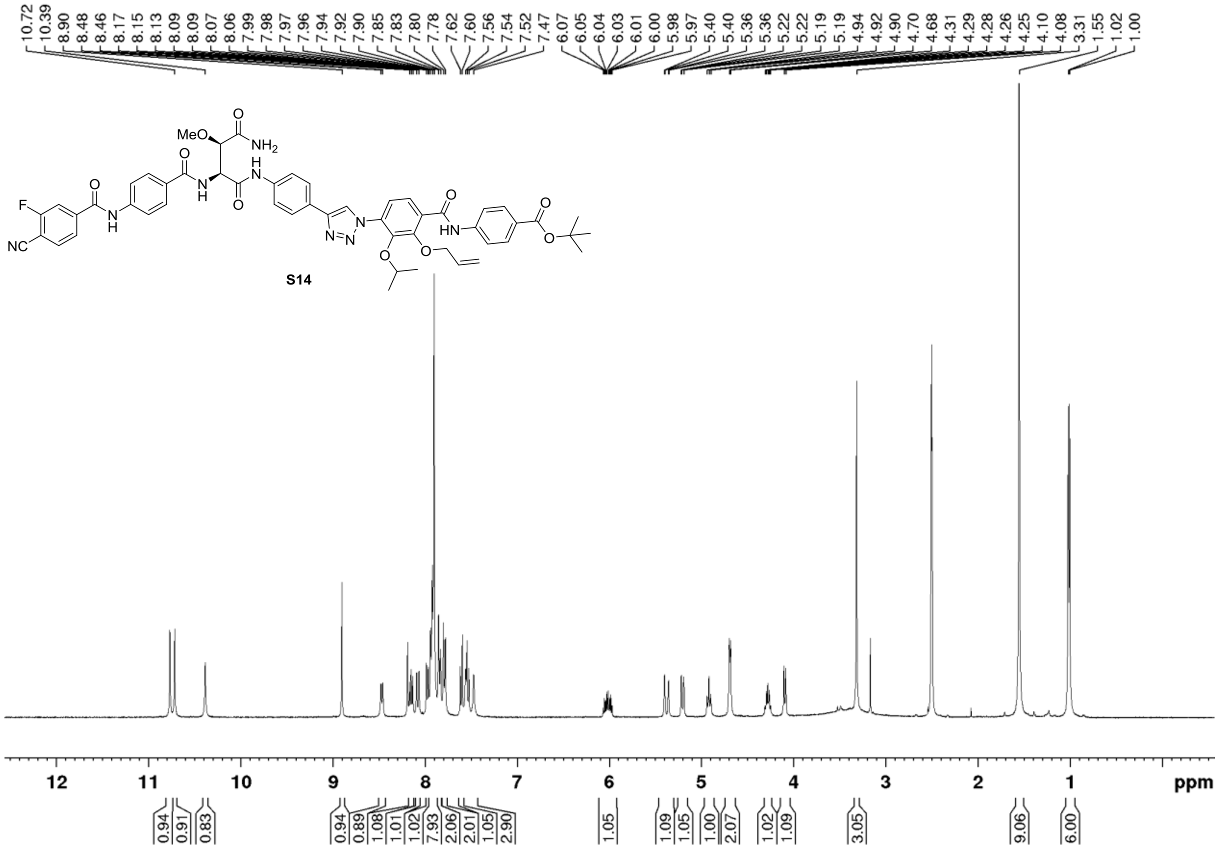
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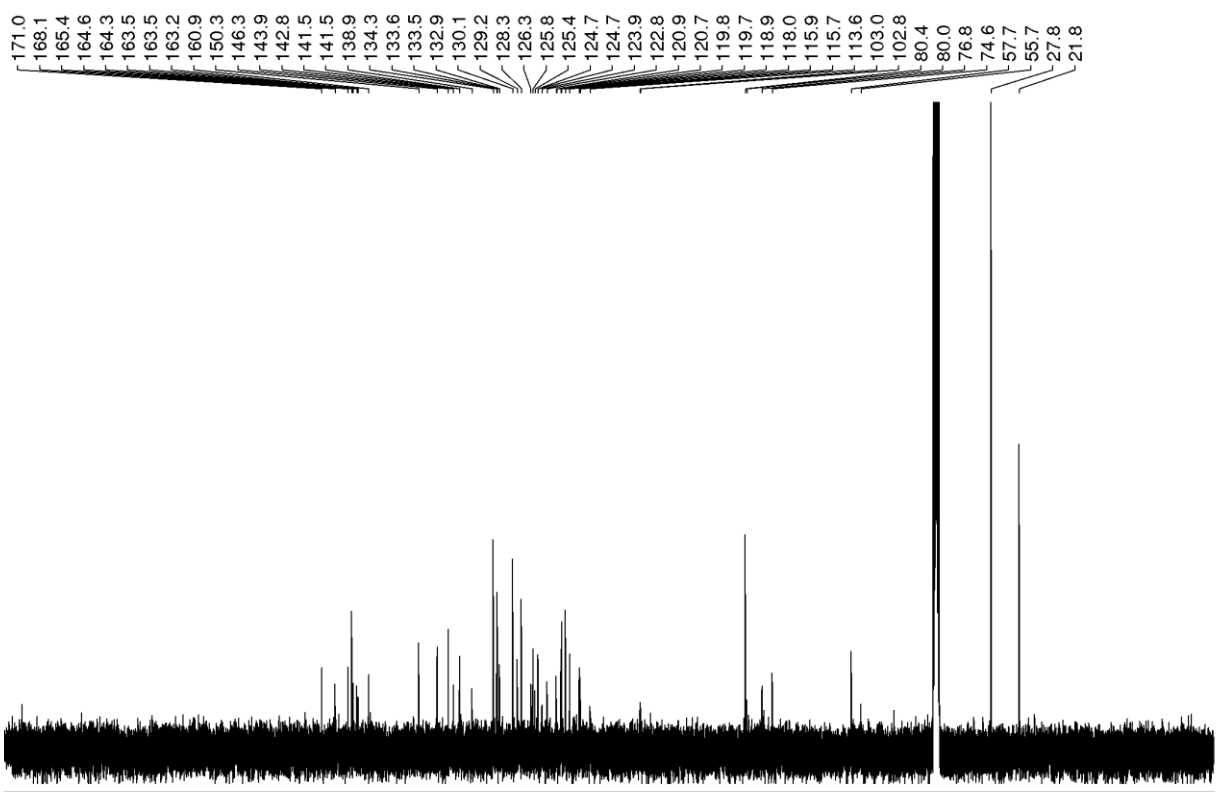
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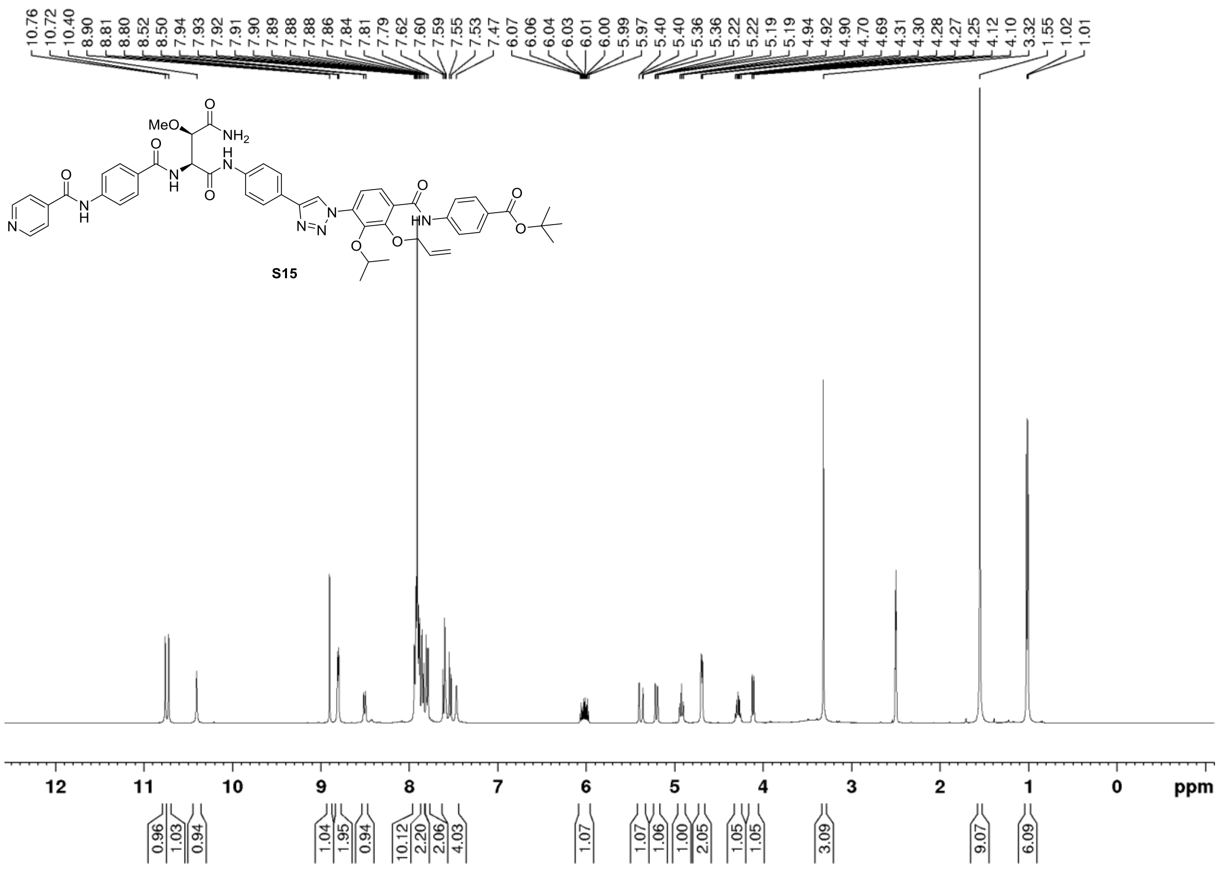
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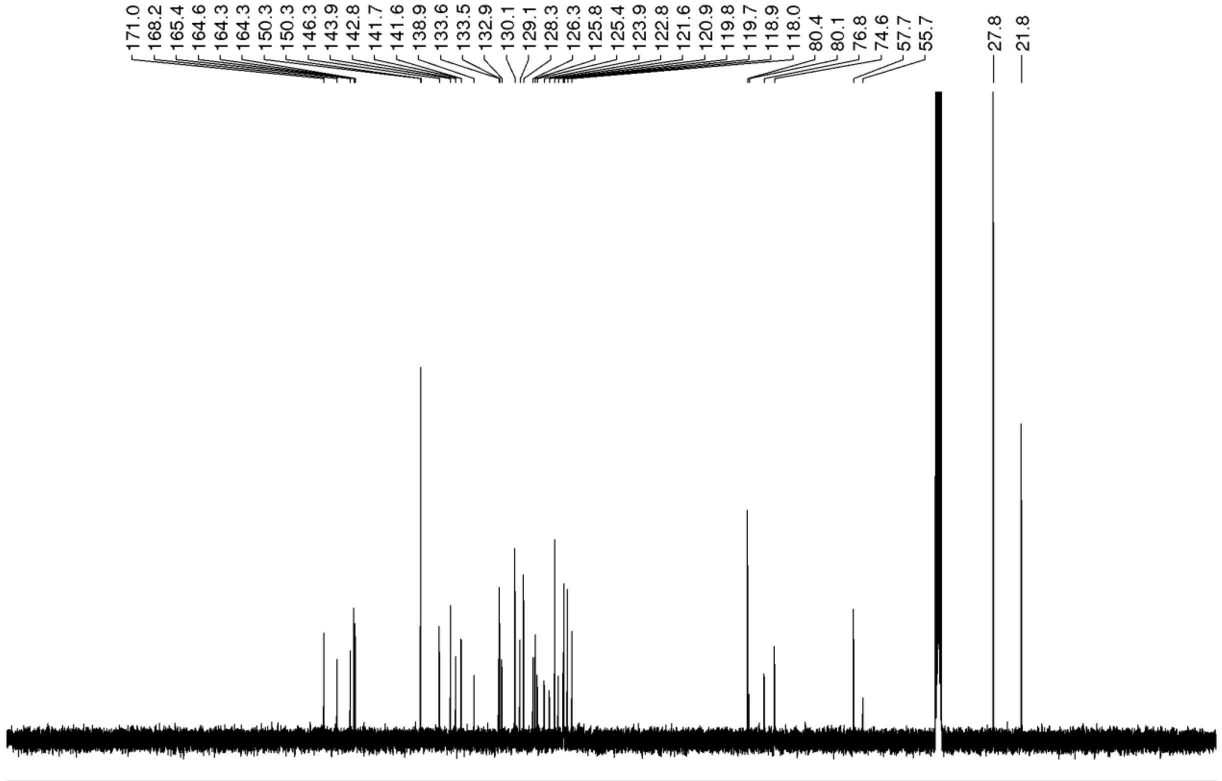
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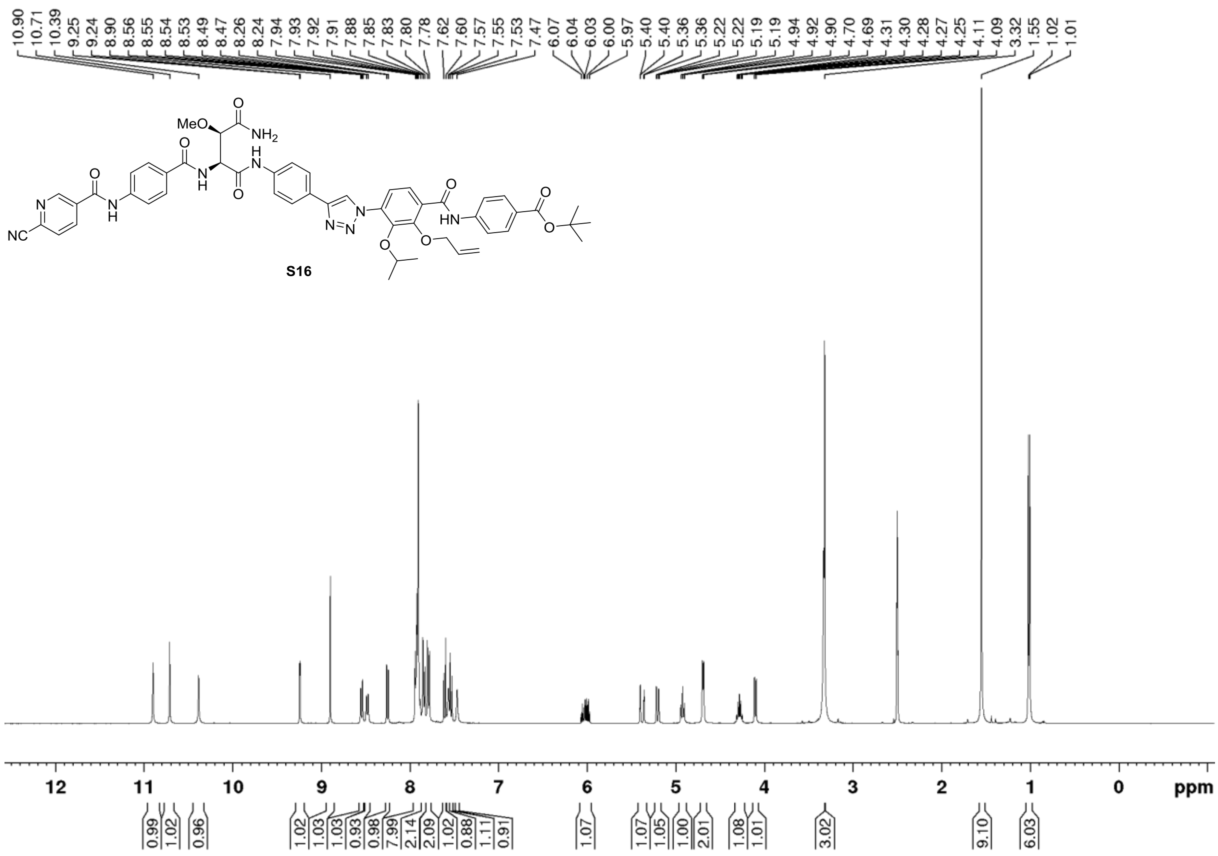
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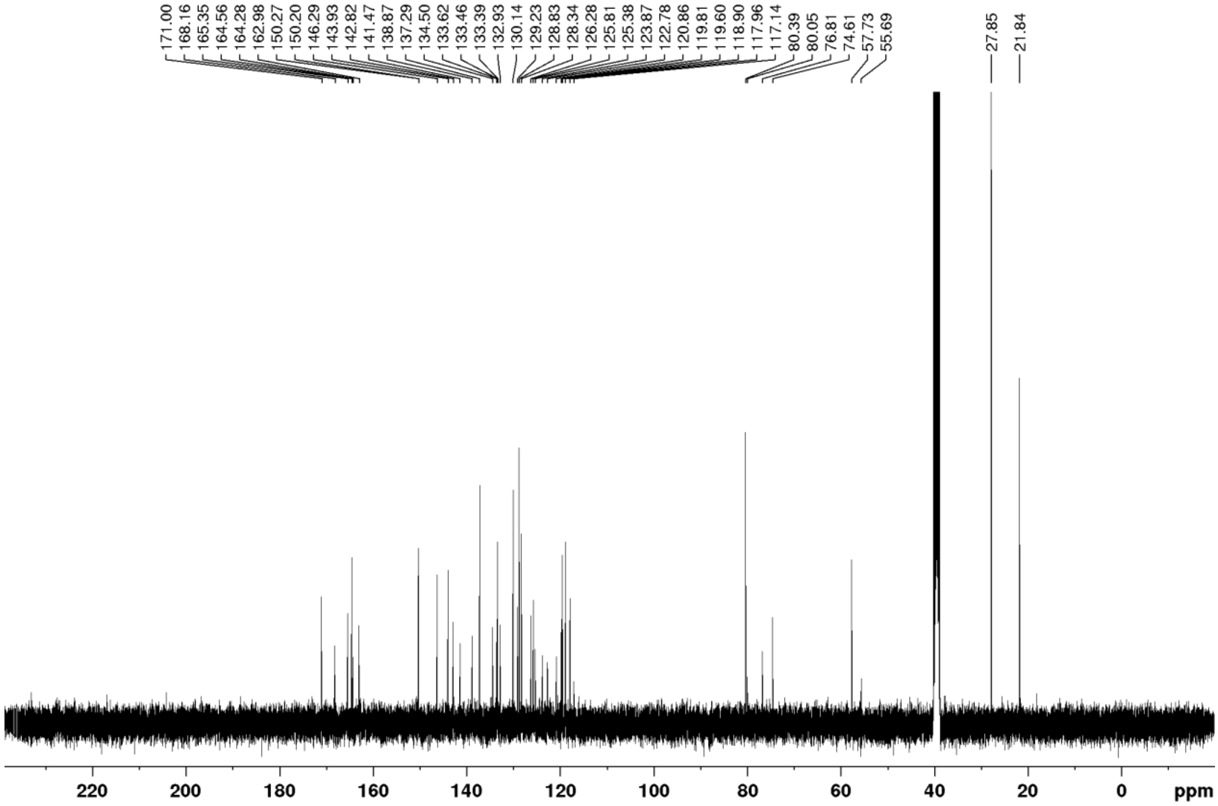
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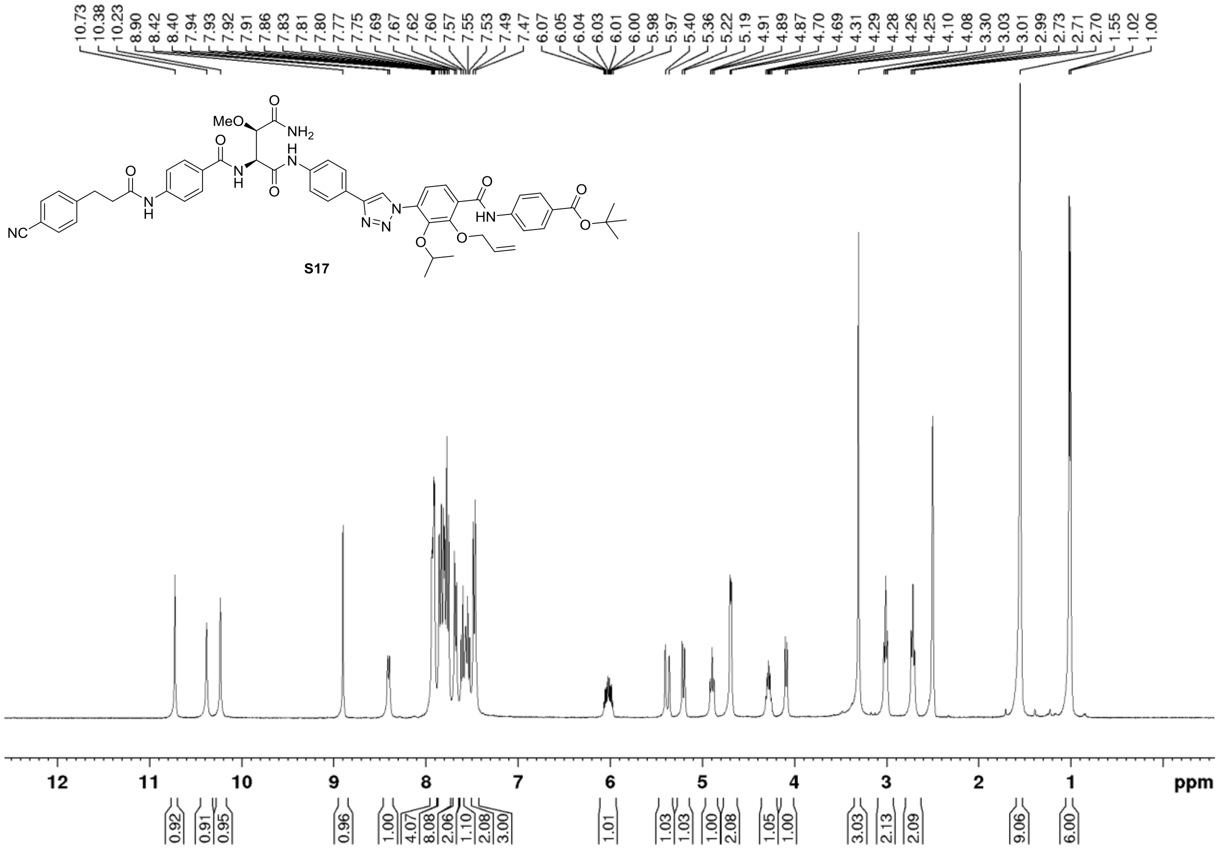
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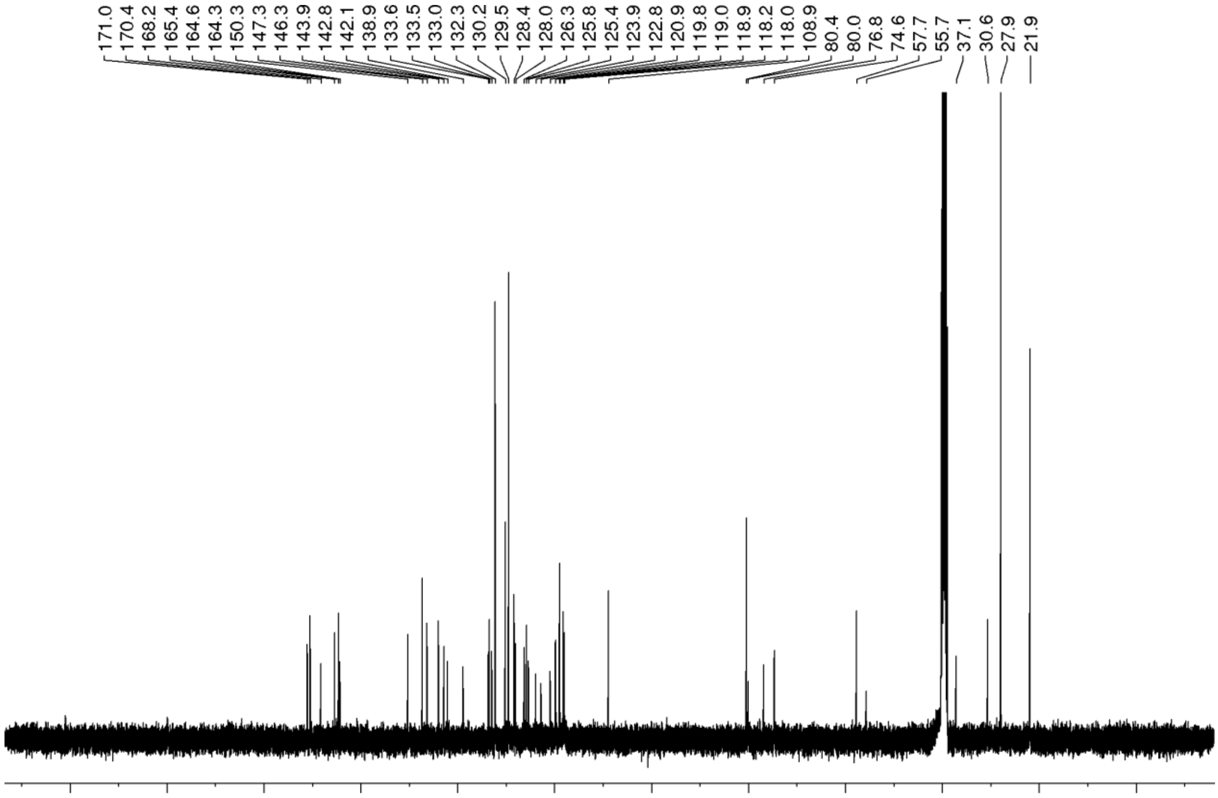
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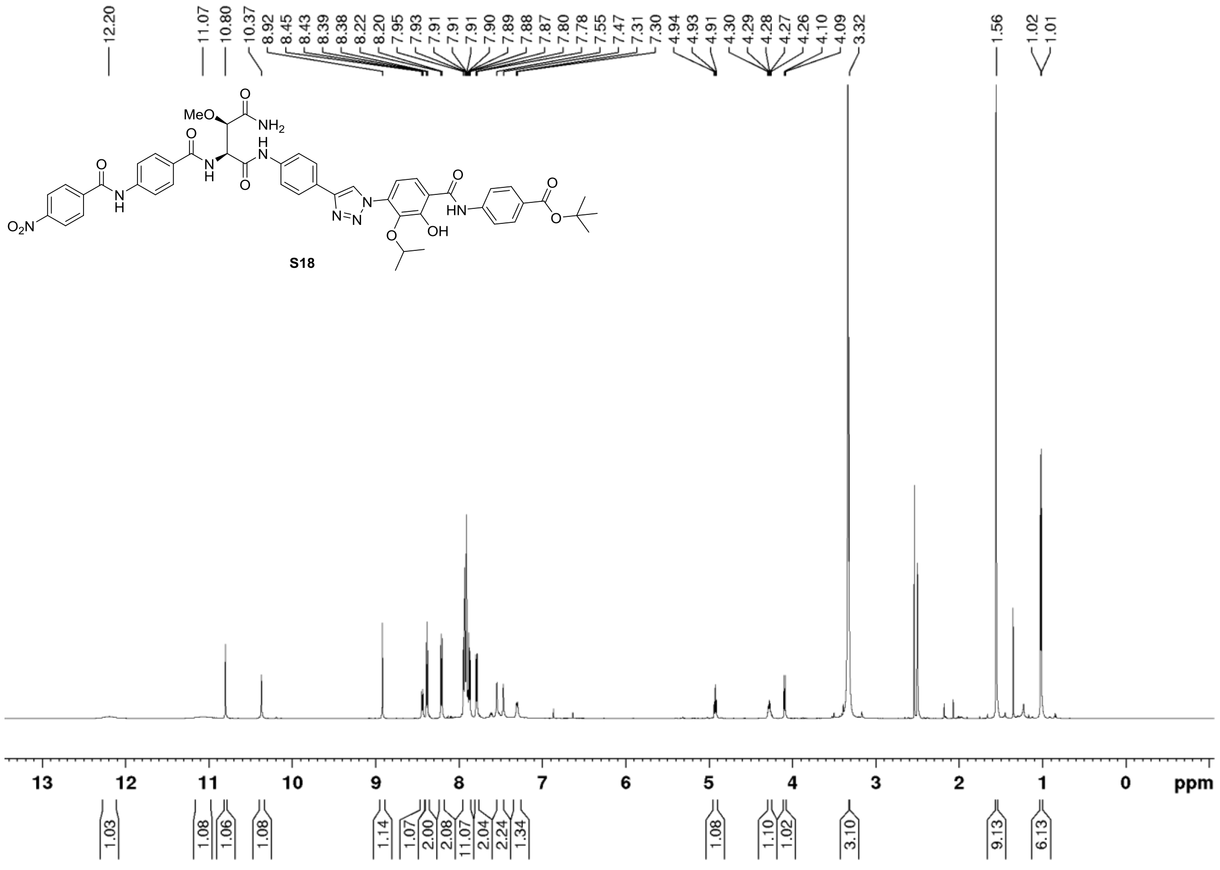
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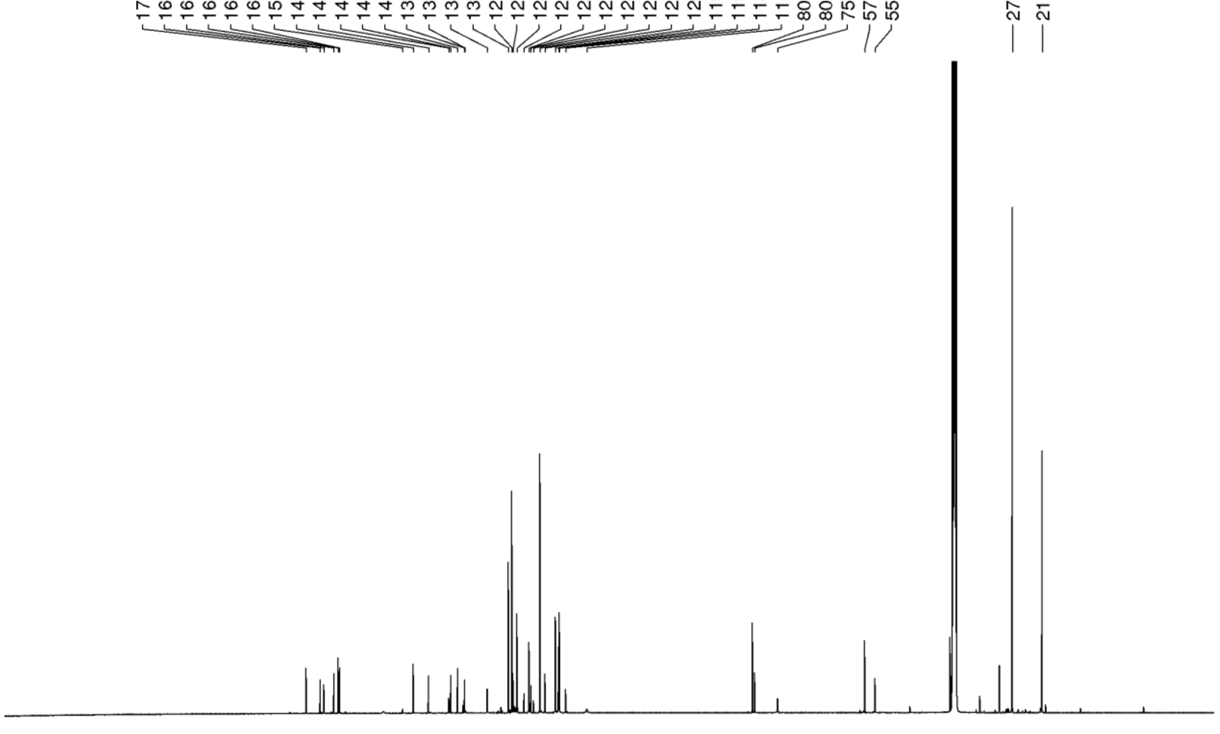
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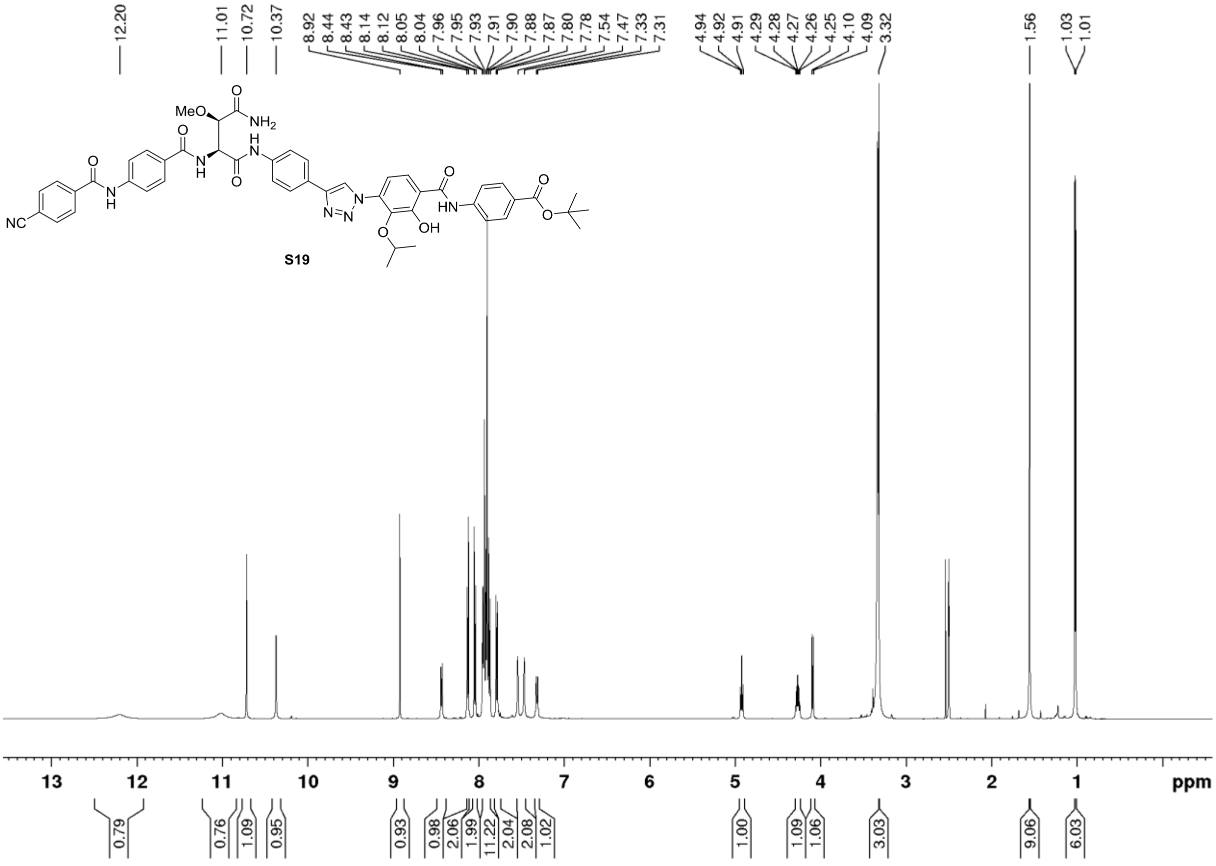
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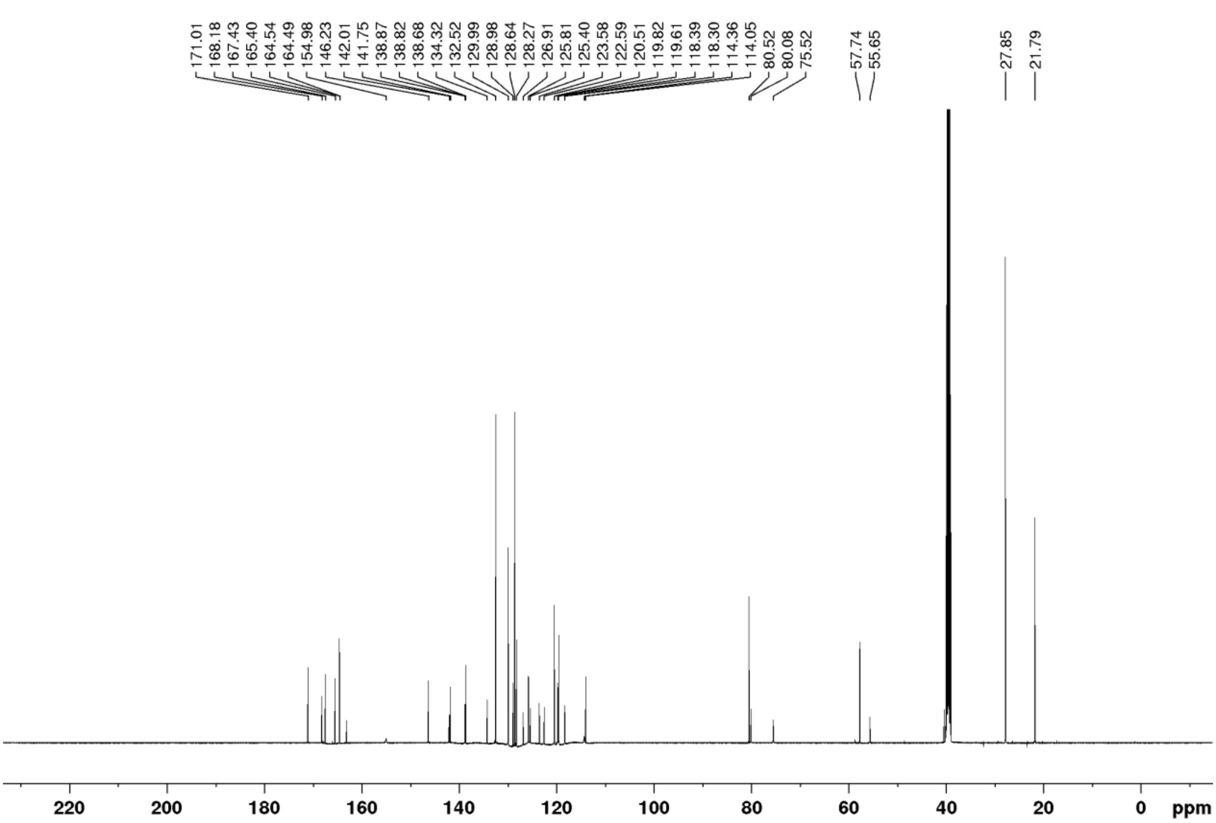
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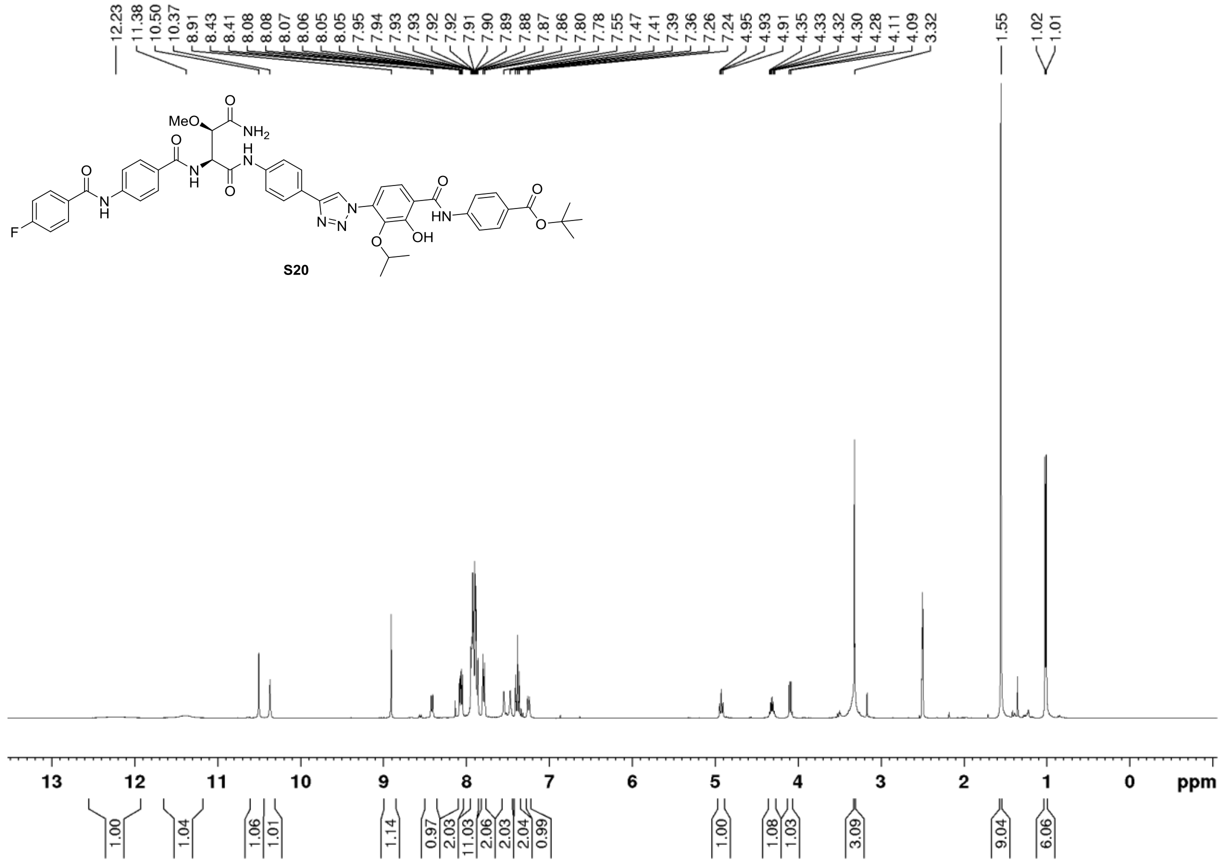
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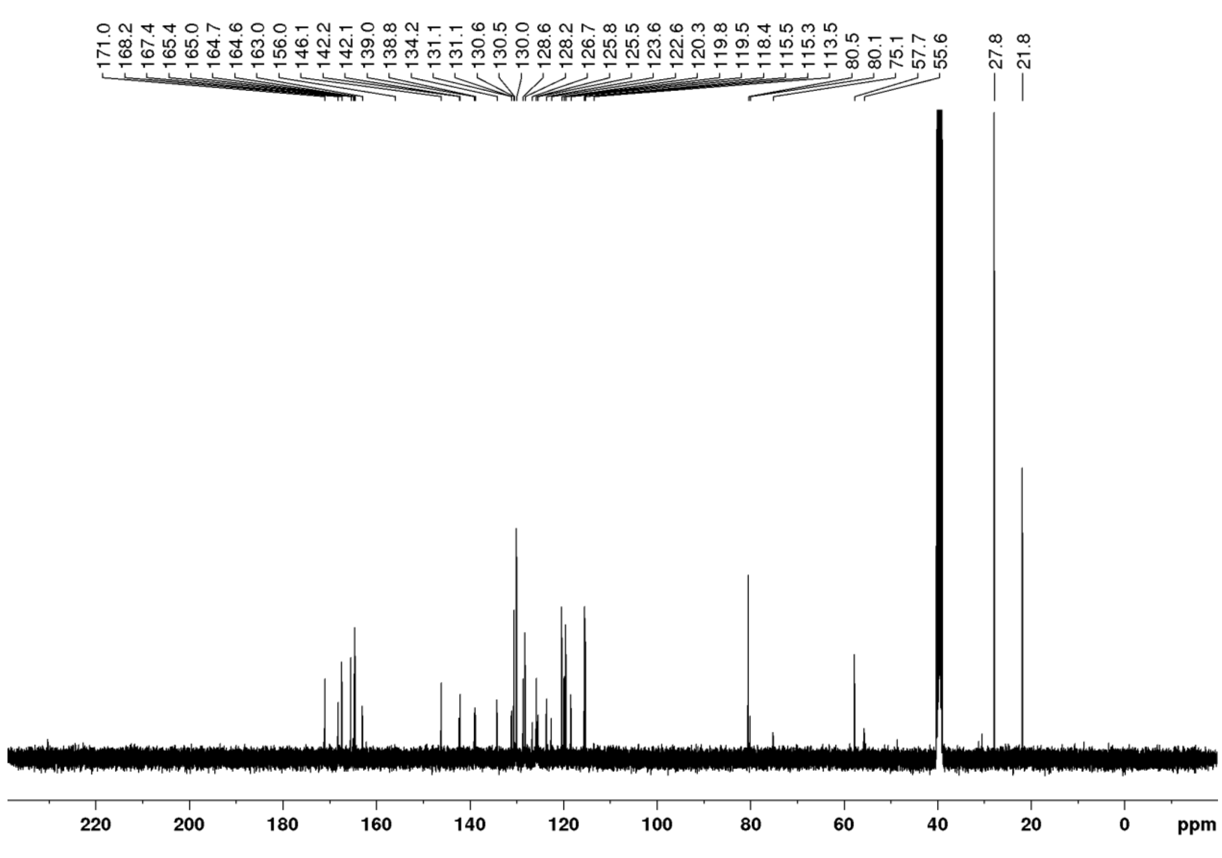
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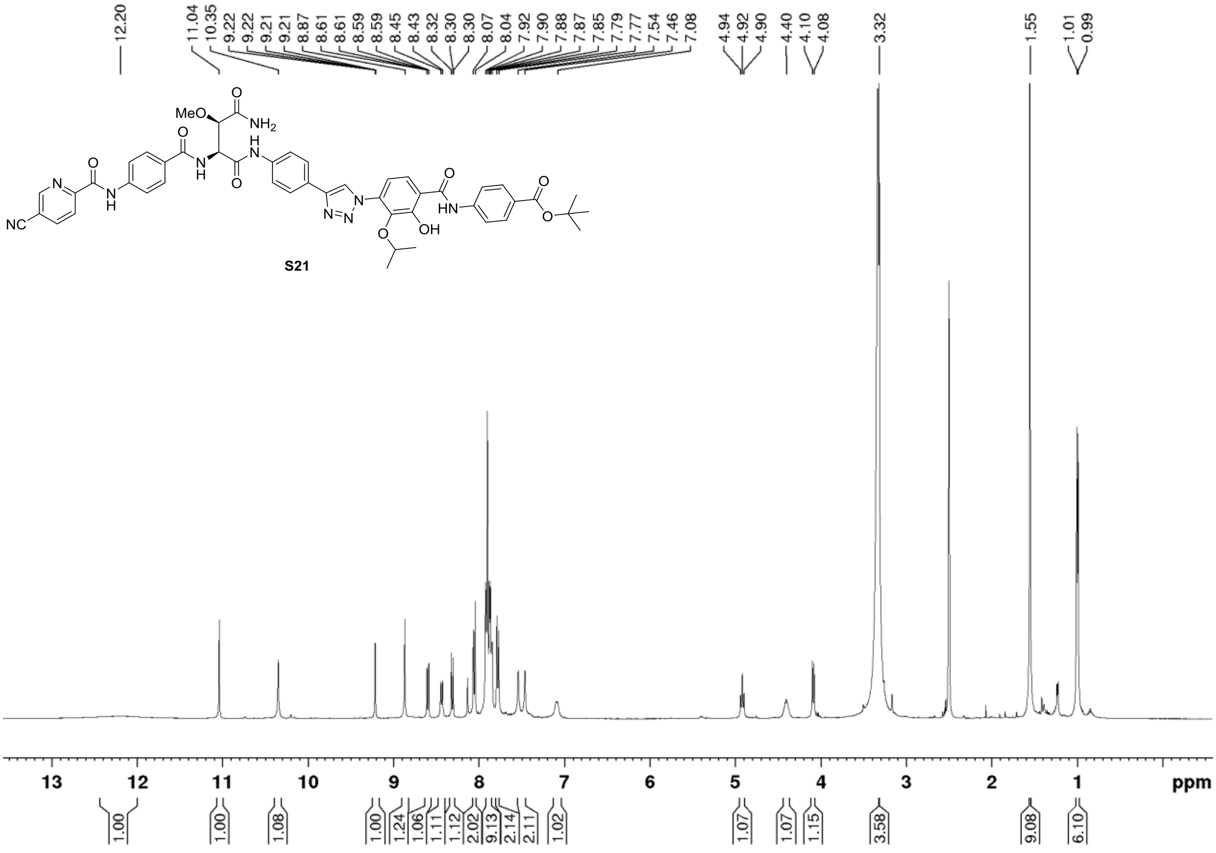
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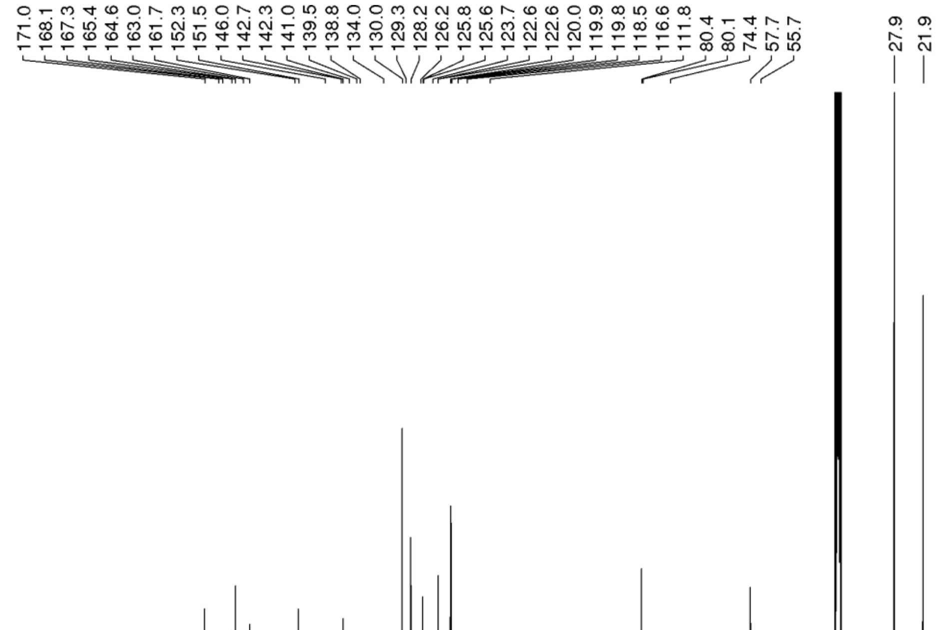
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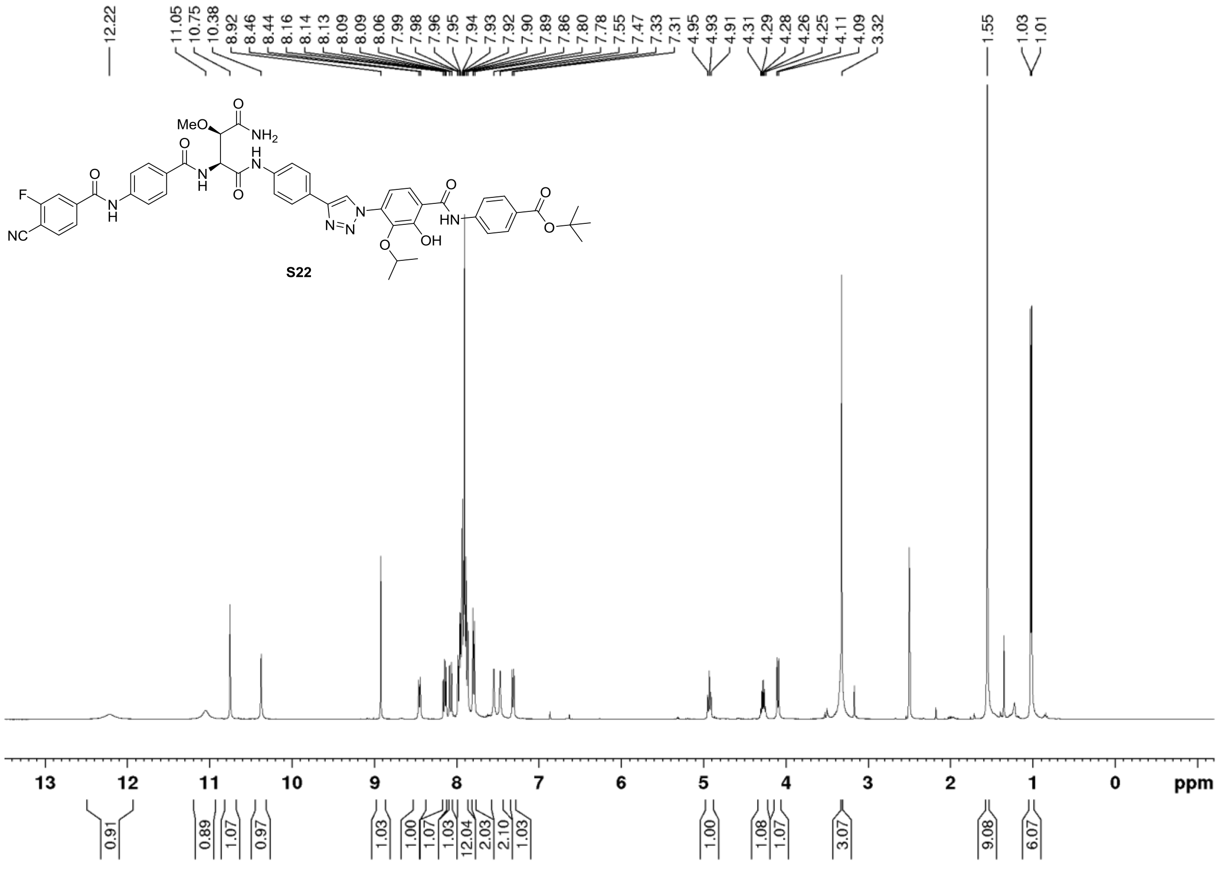
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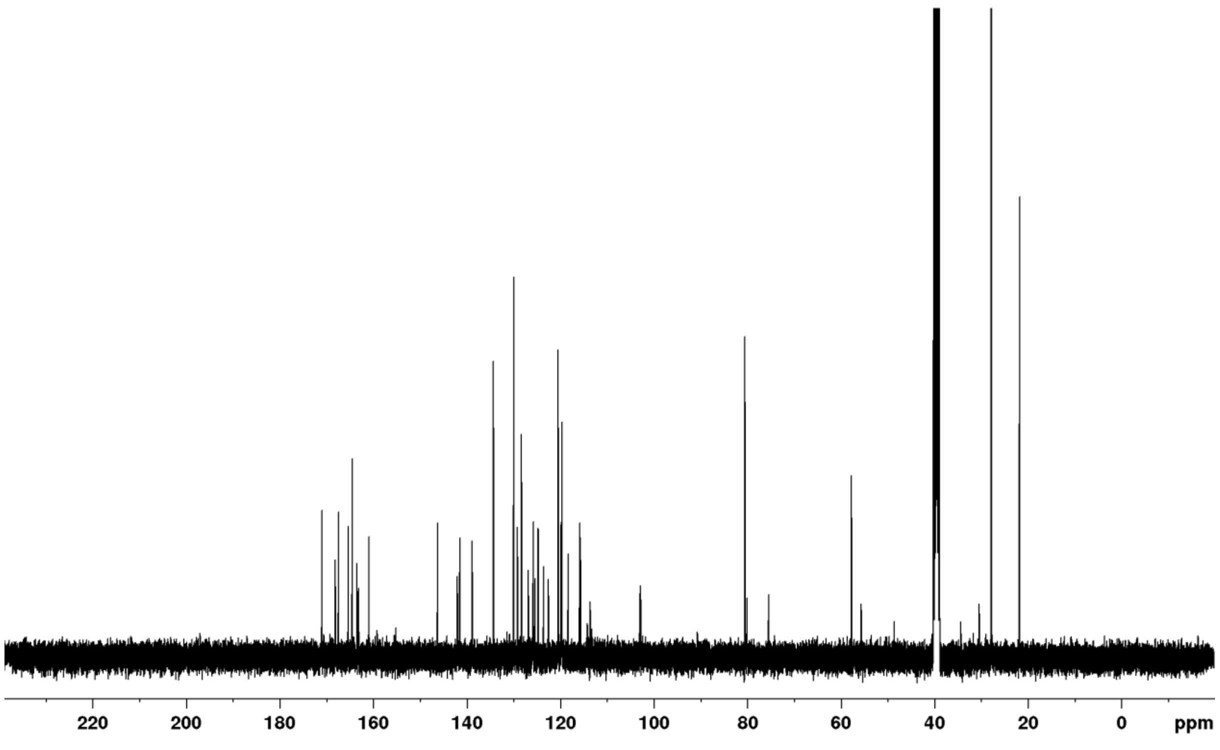
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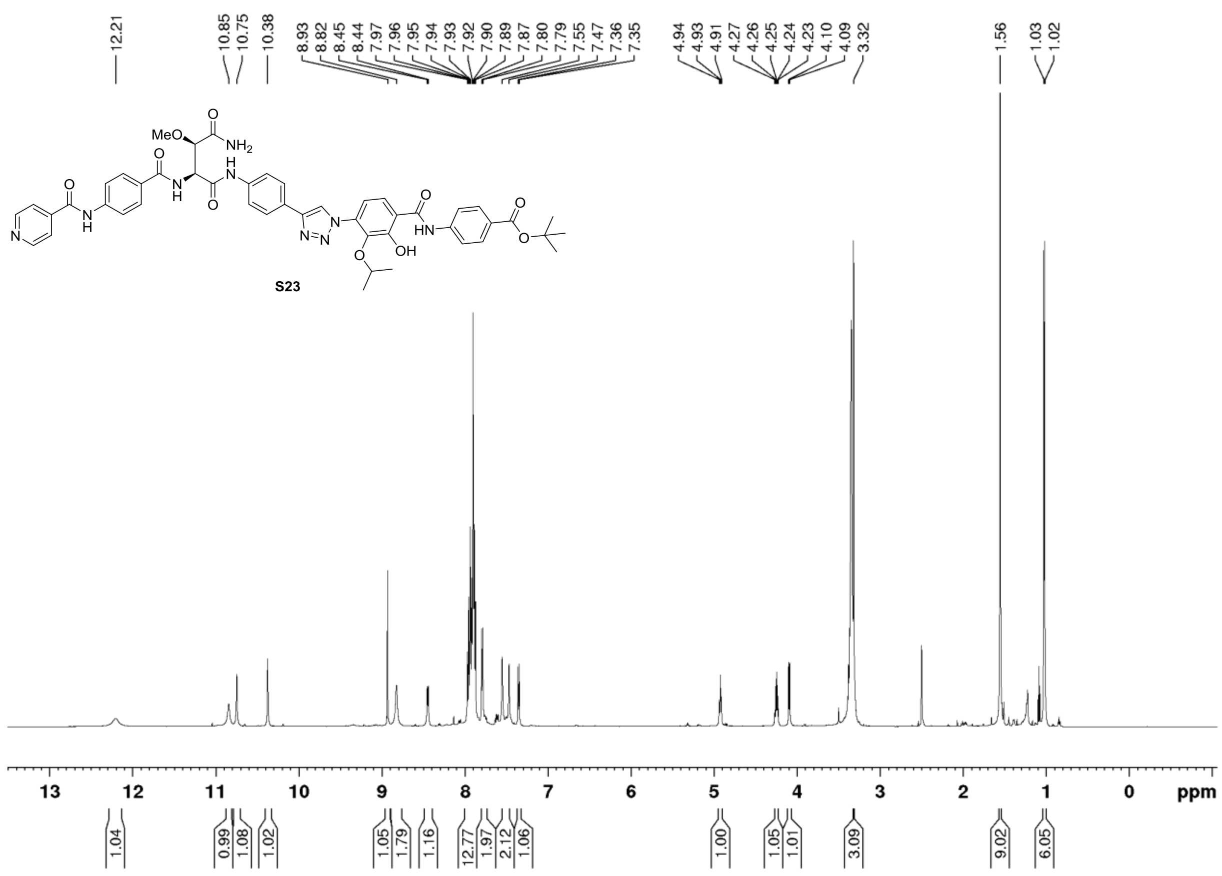

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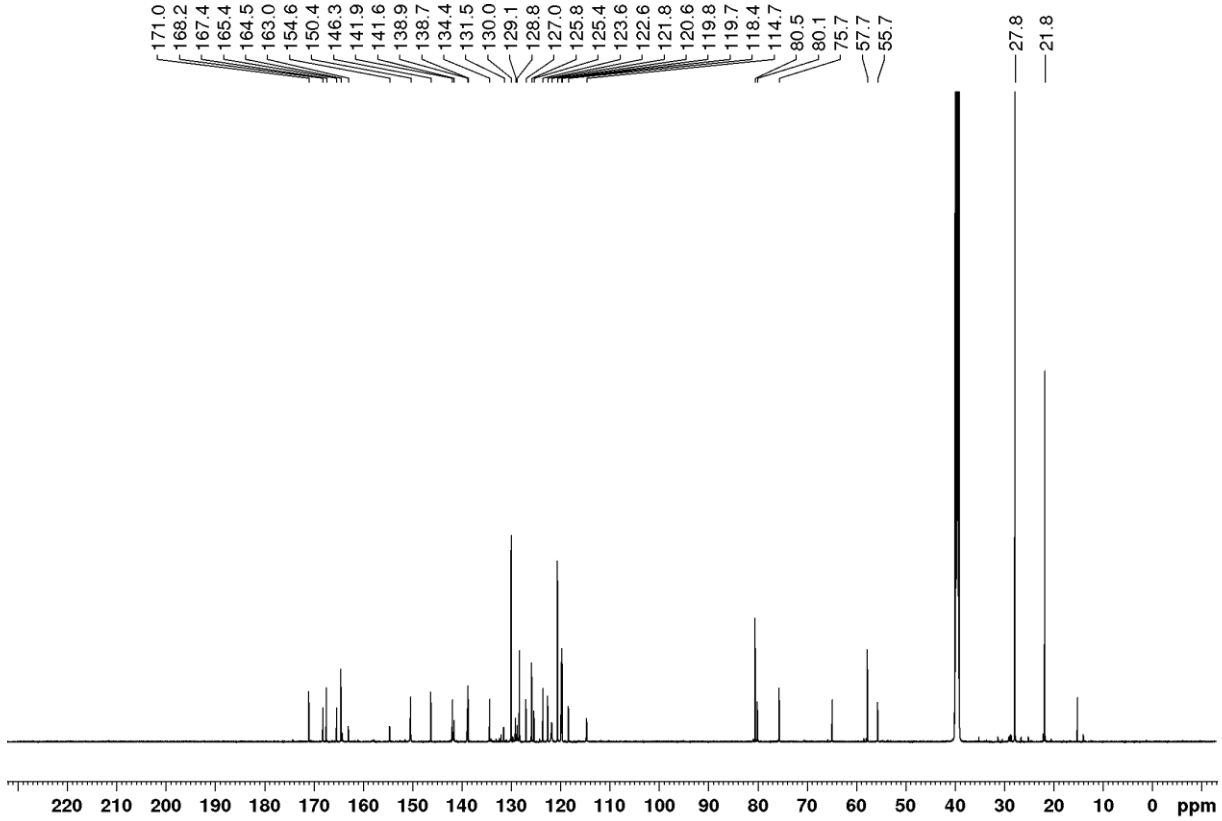

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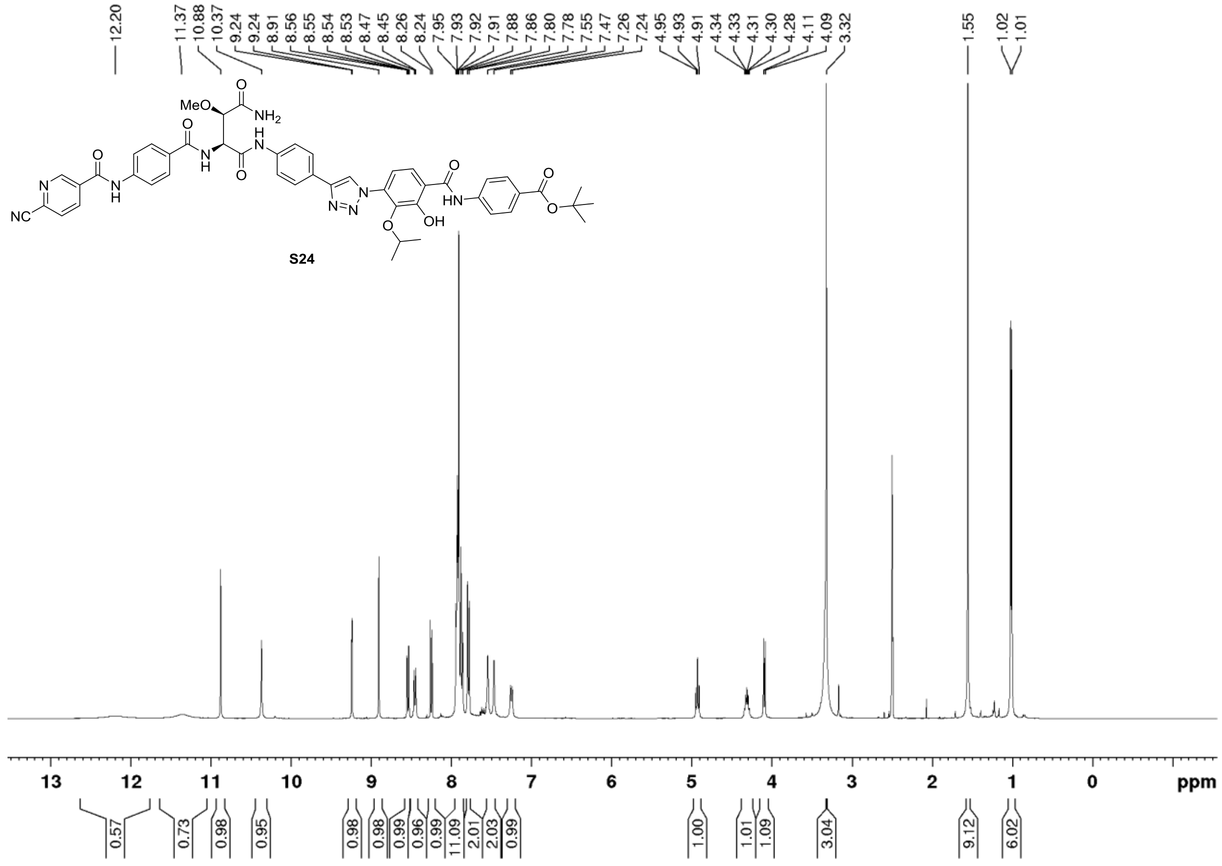
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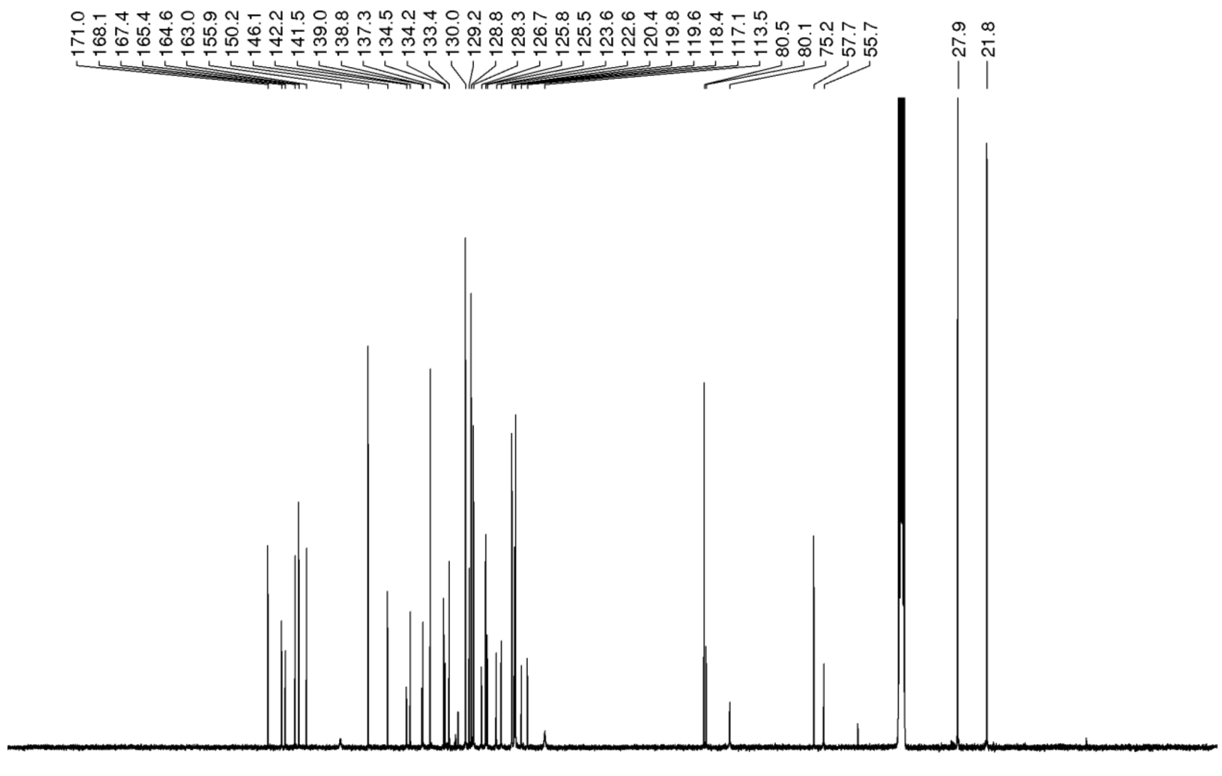
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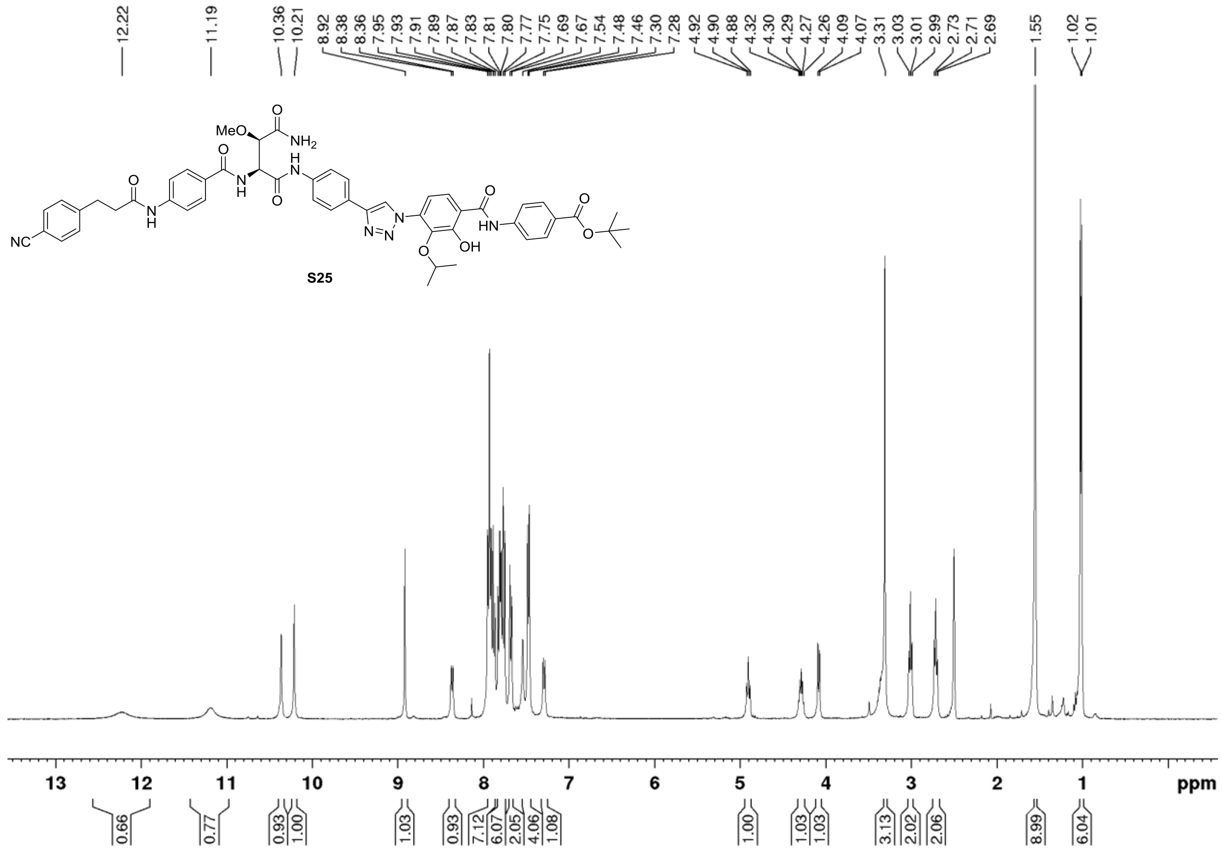
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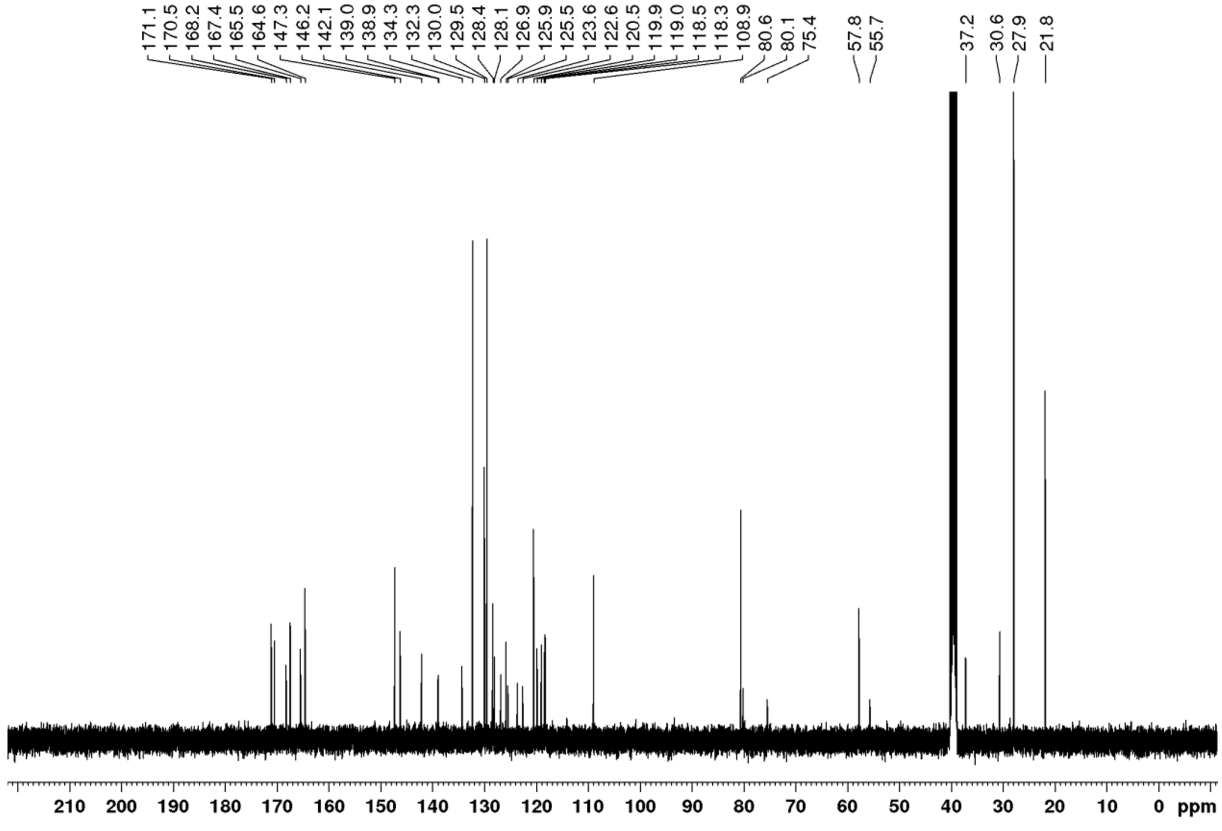

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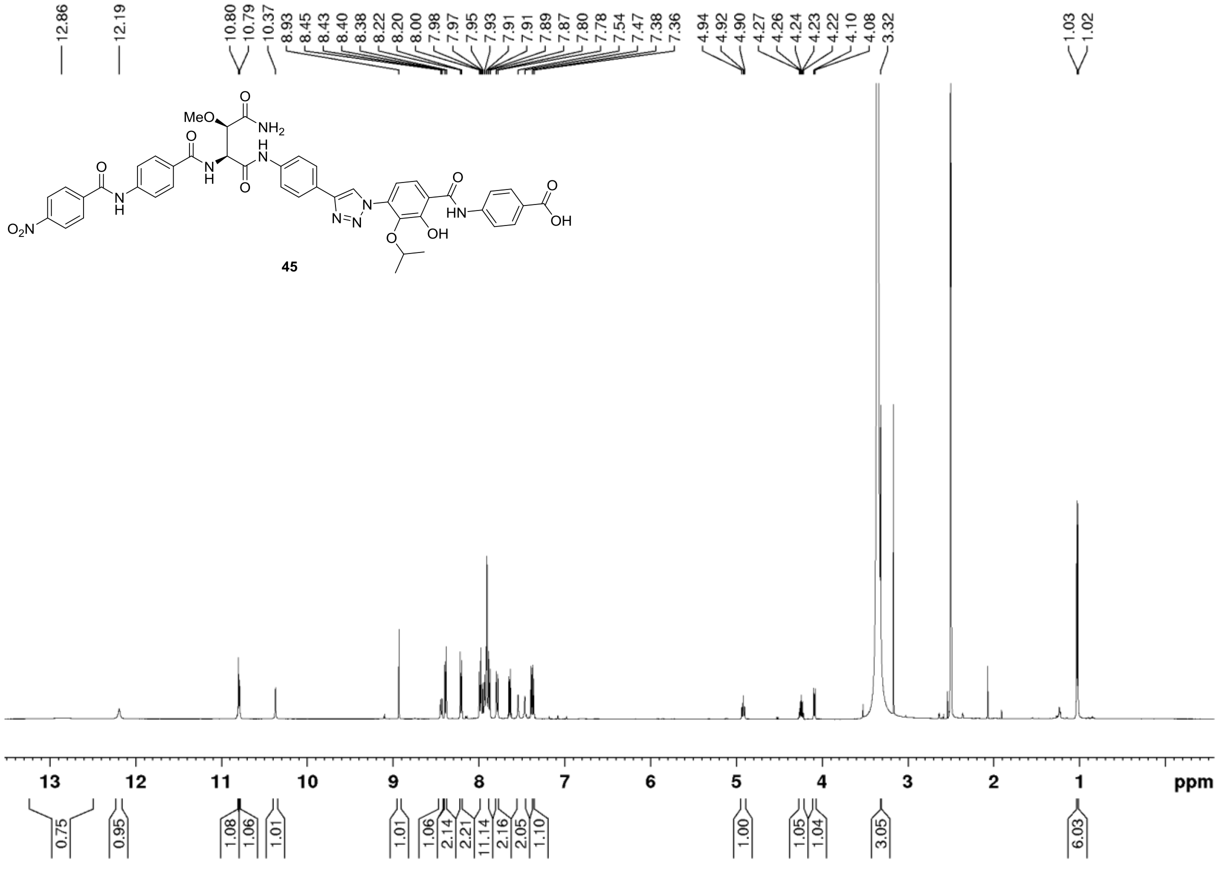
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4.10
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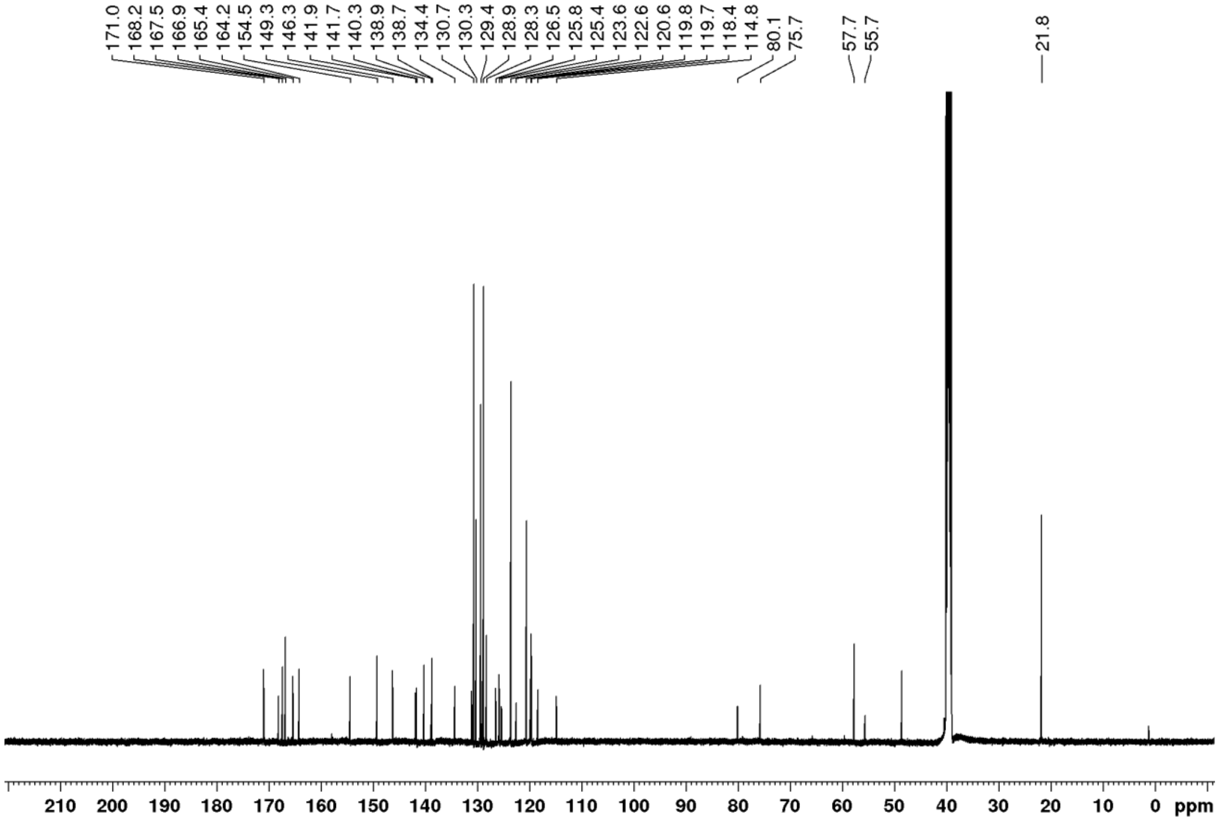
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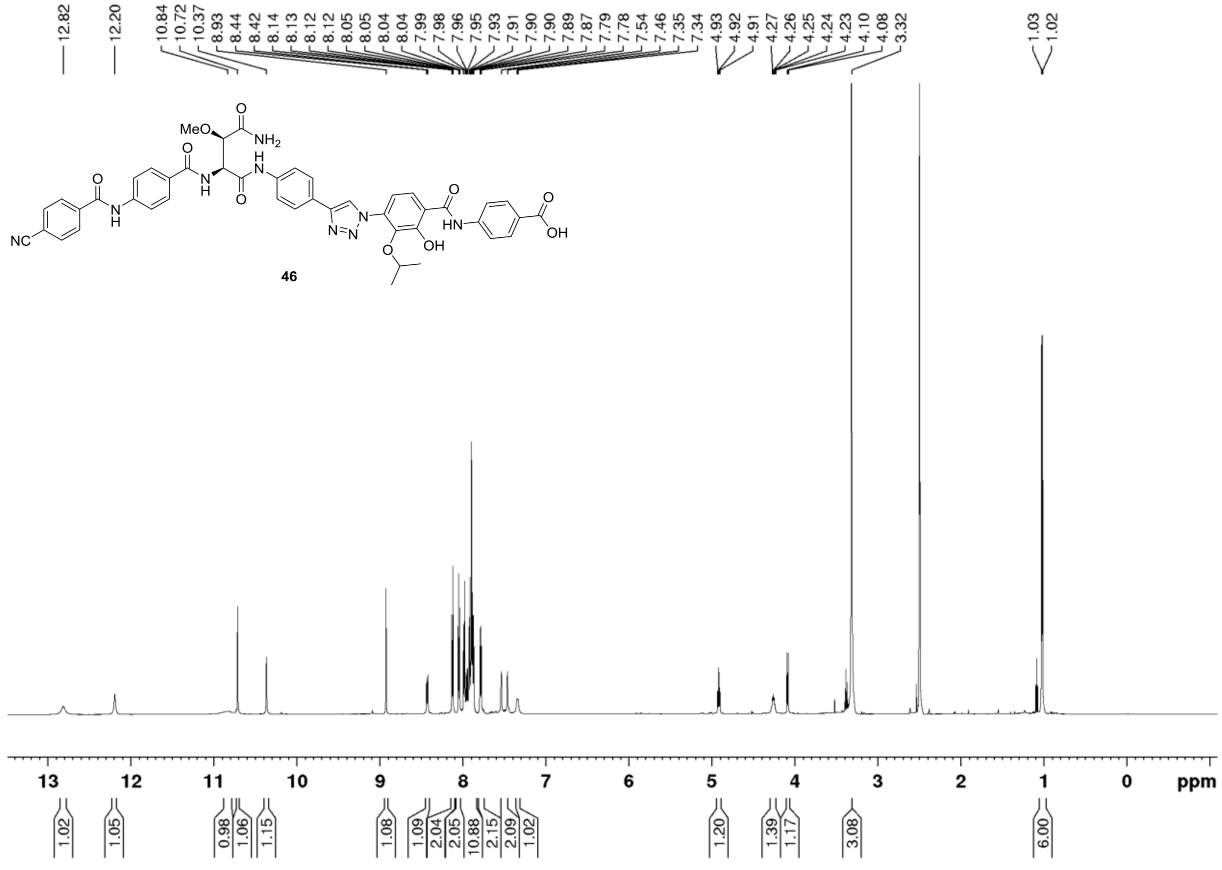
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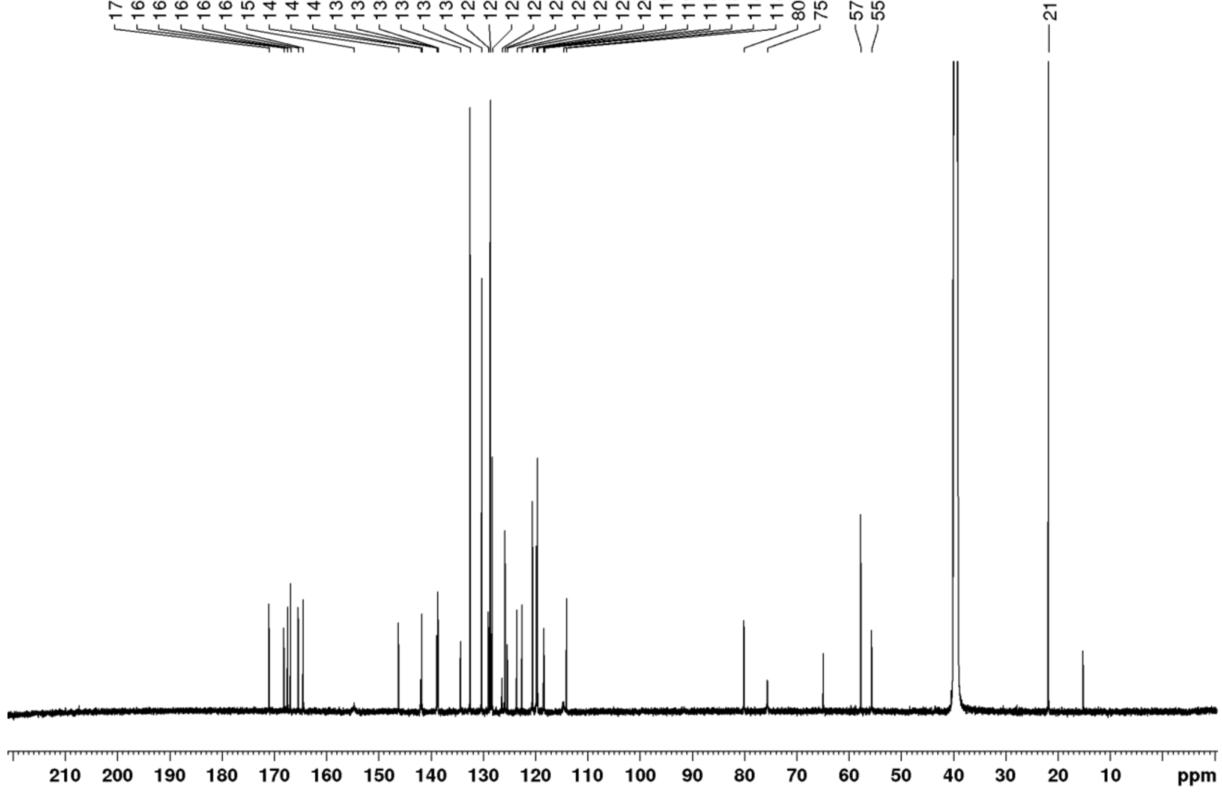
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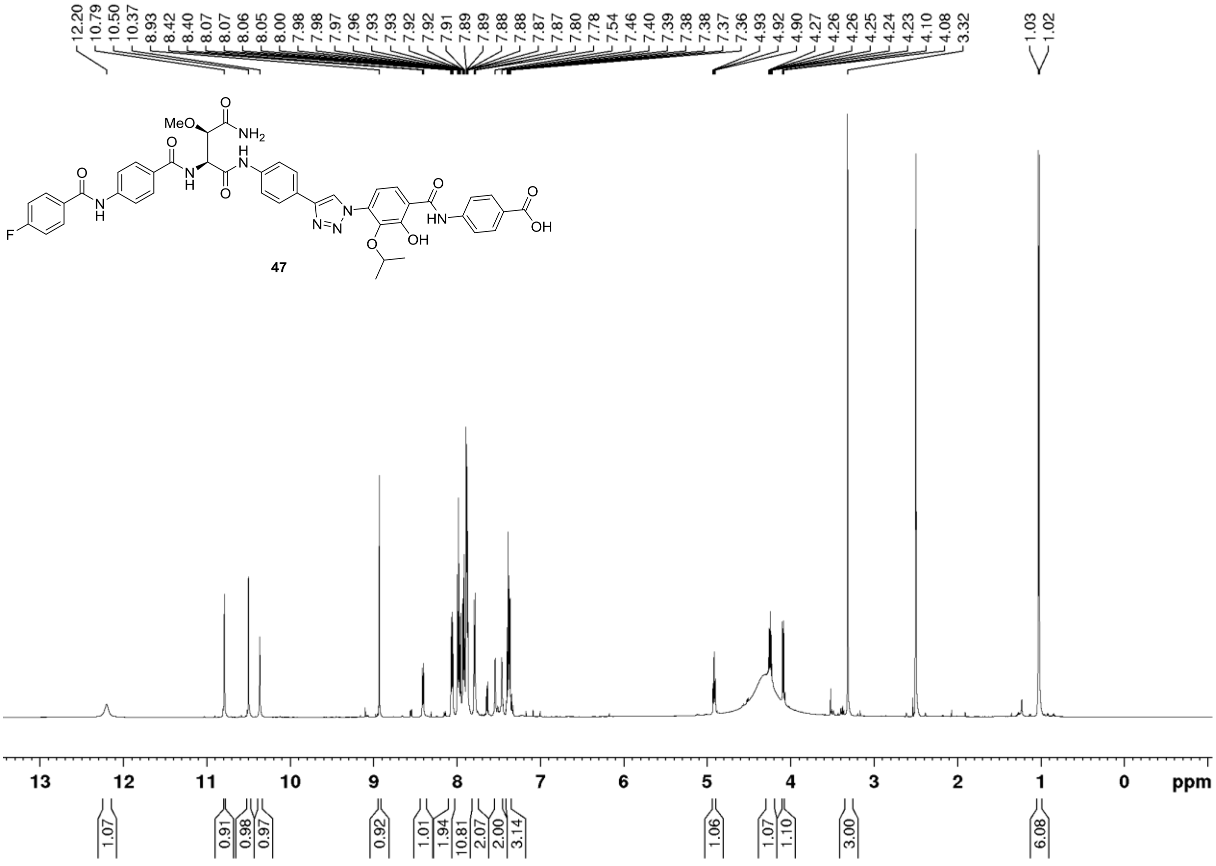
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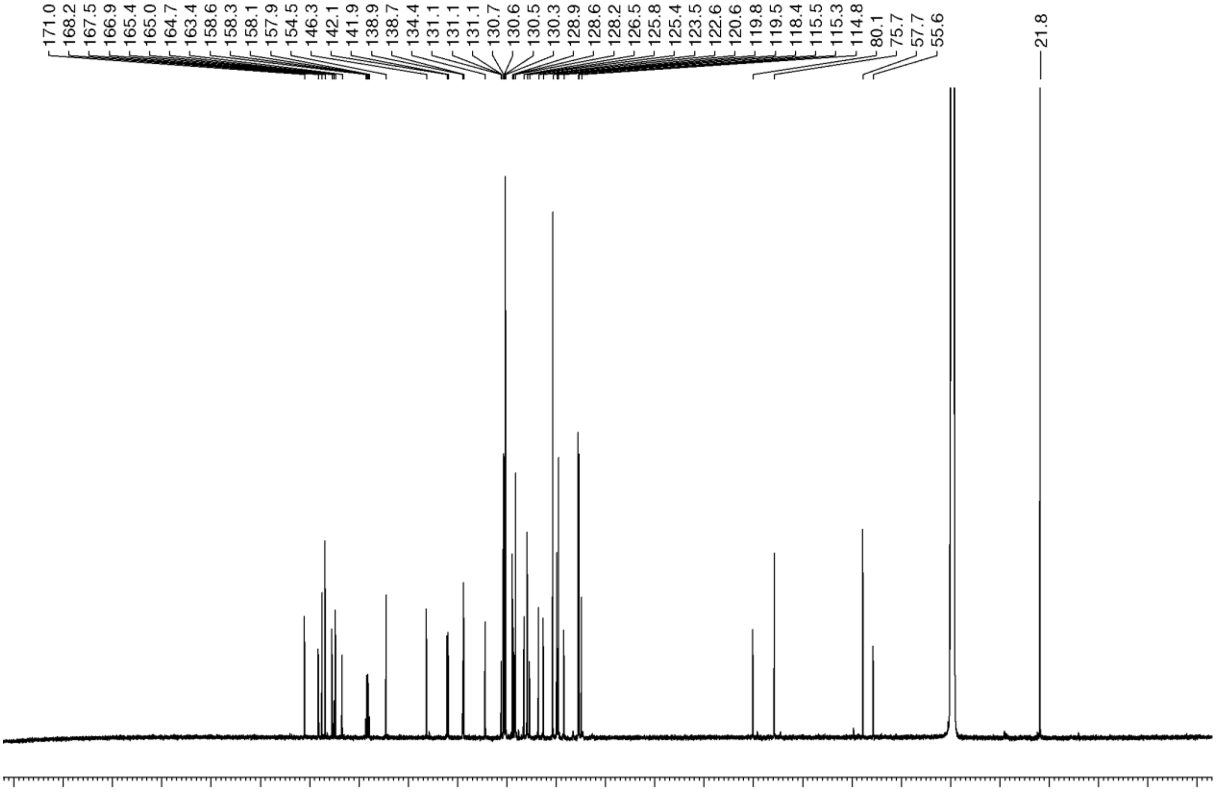
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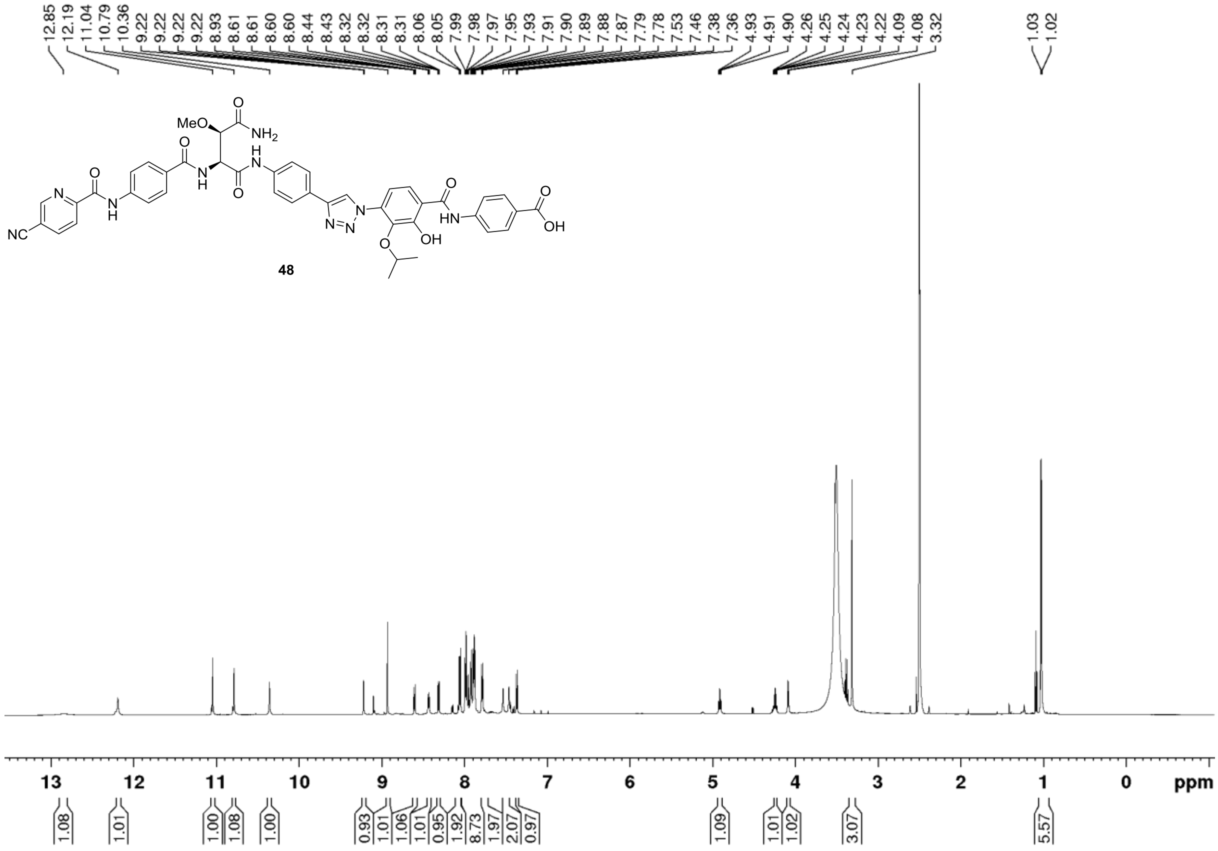
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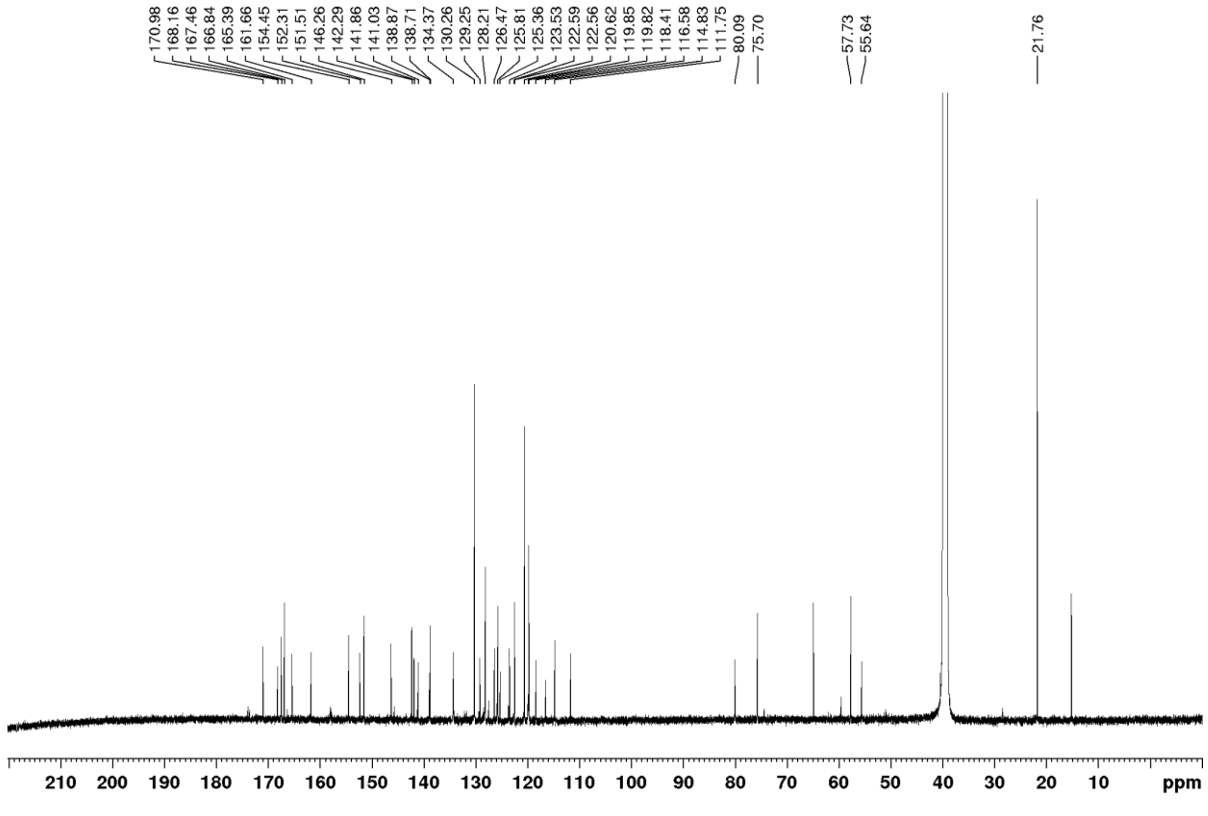
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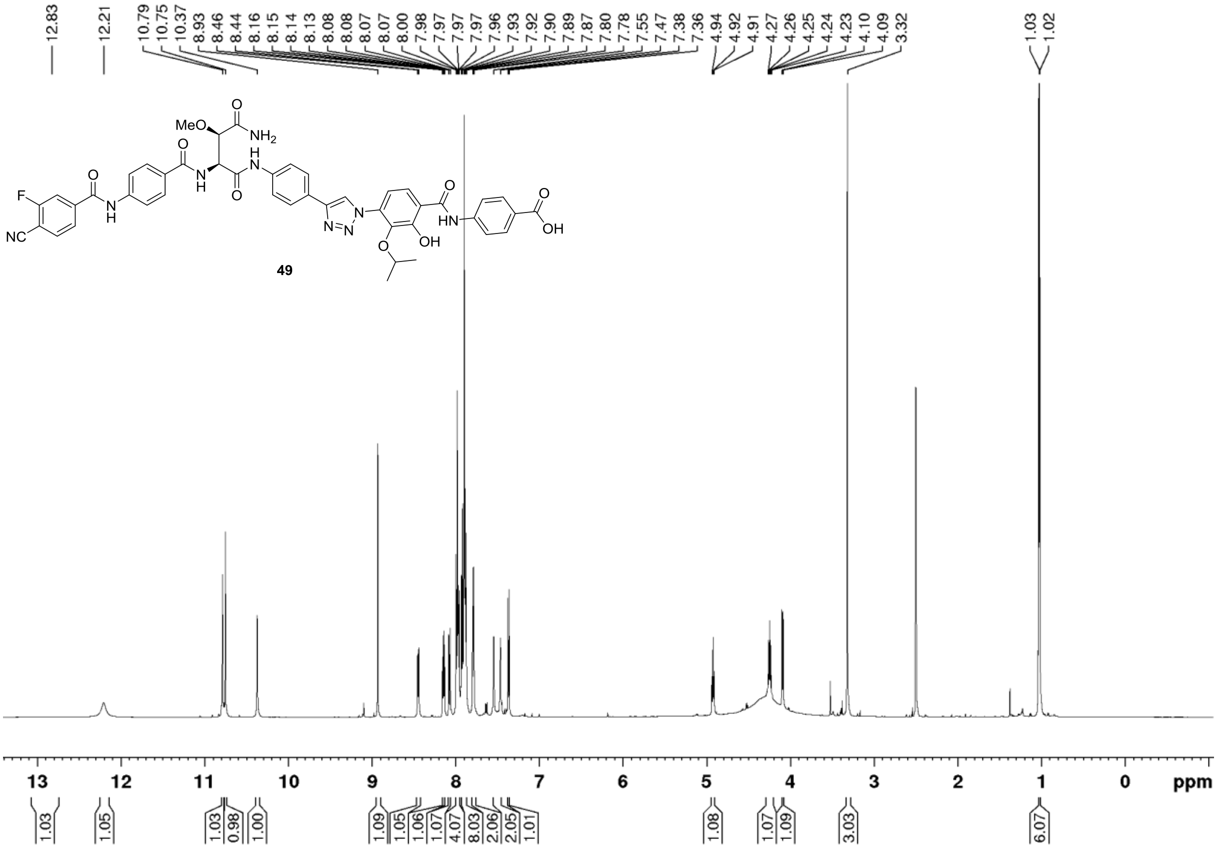

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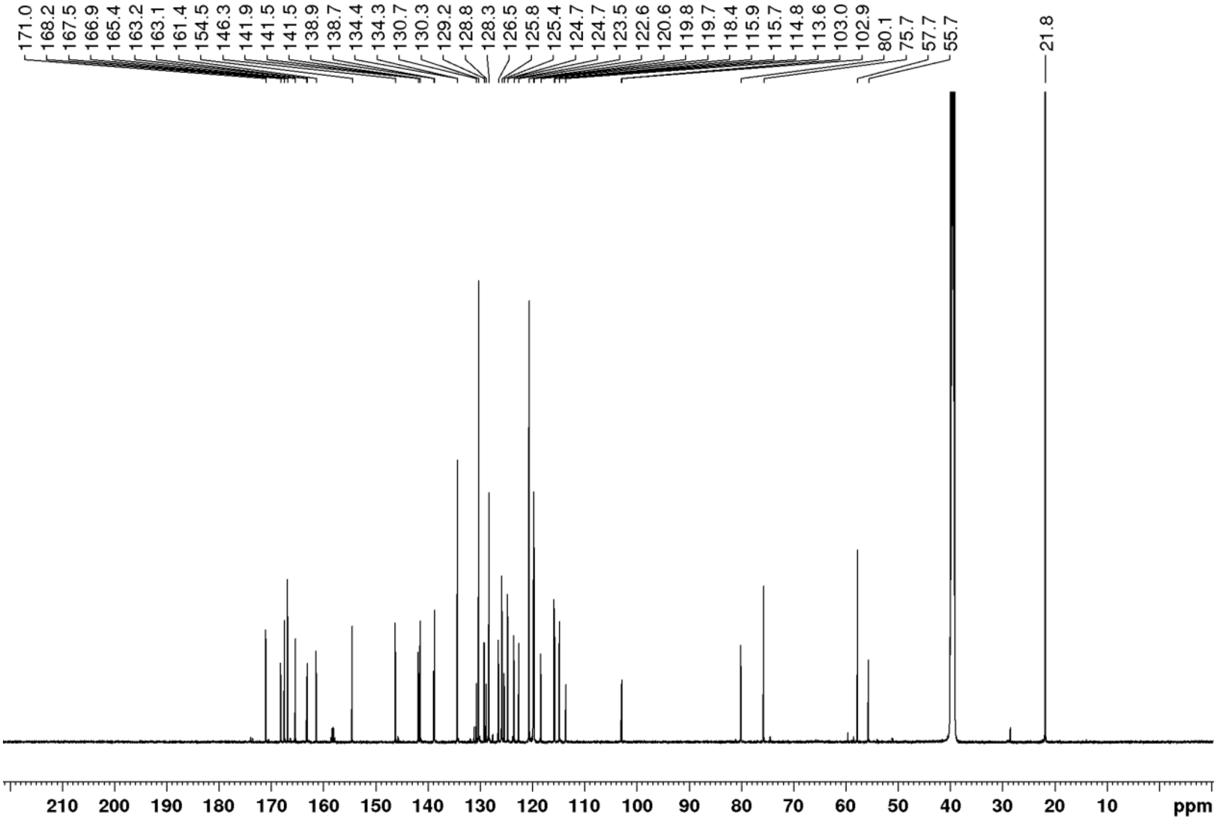

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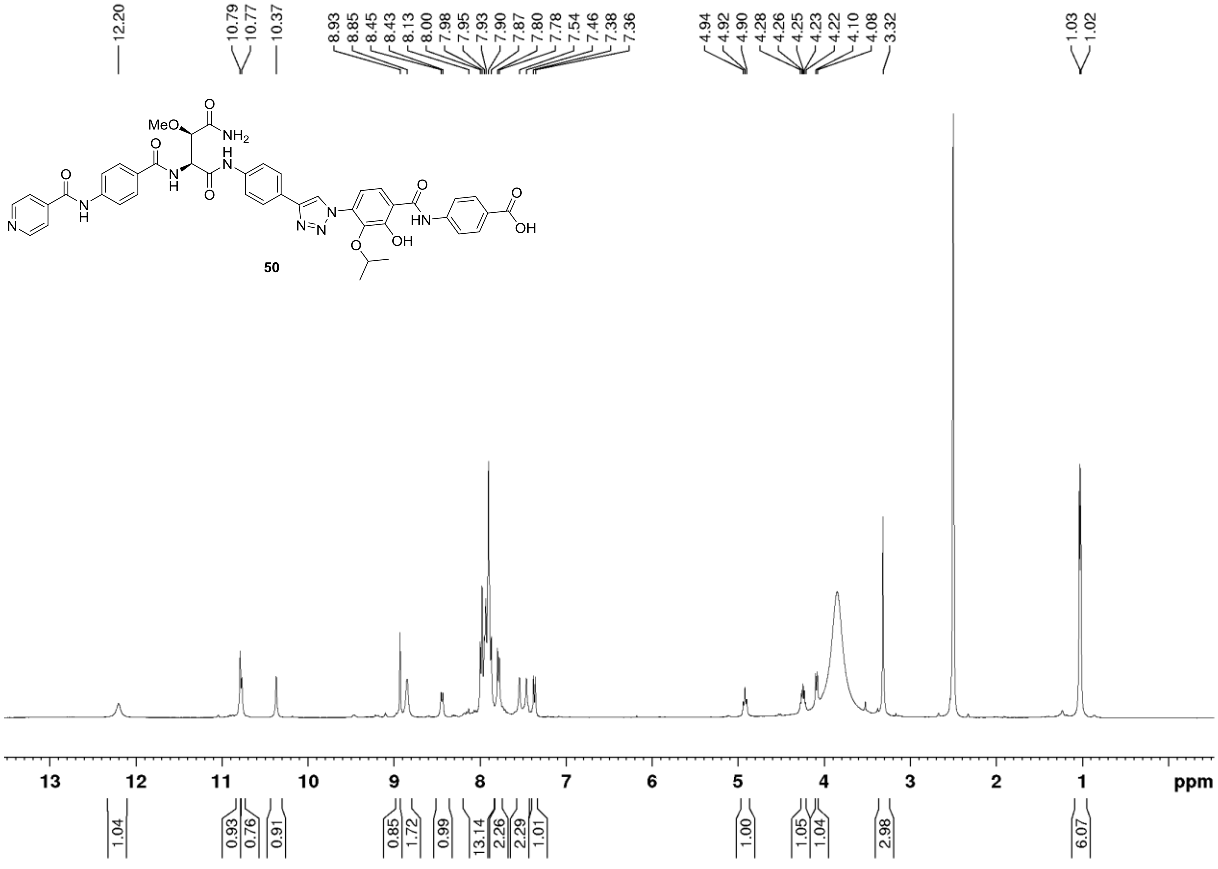12.20
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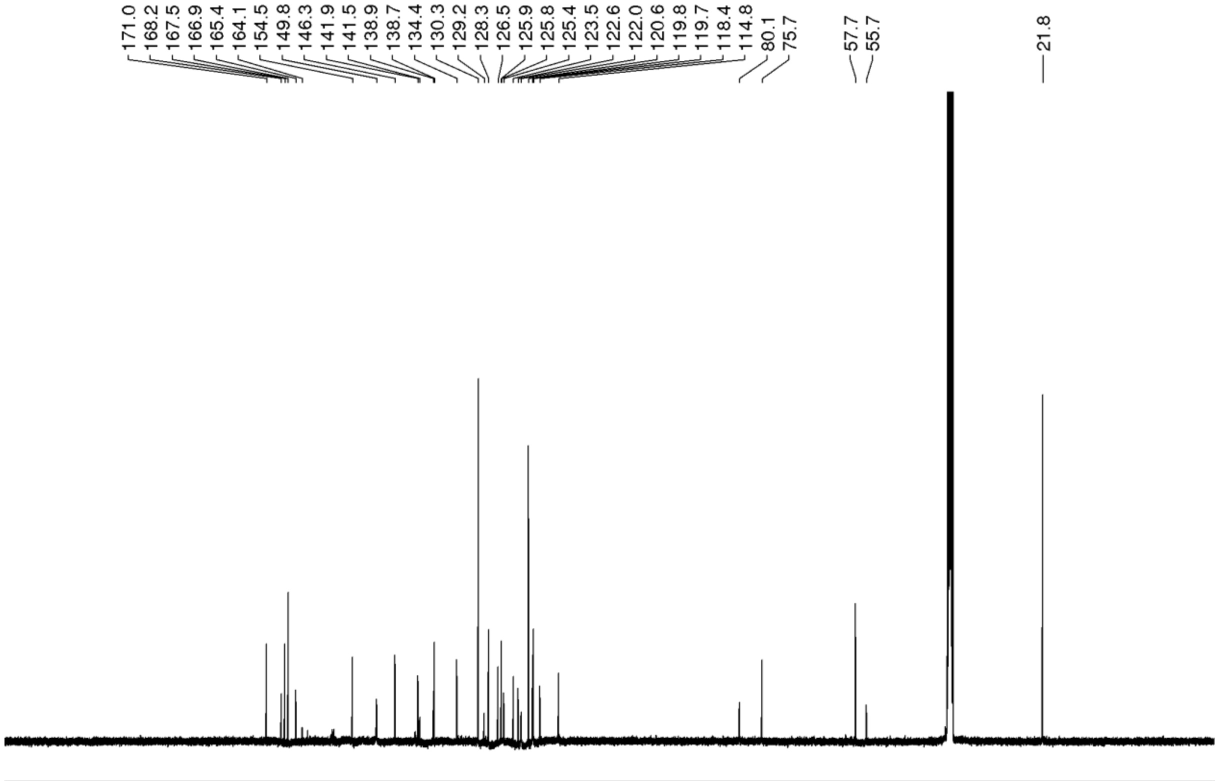171.0
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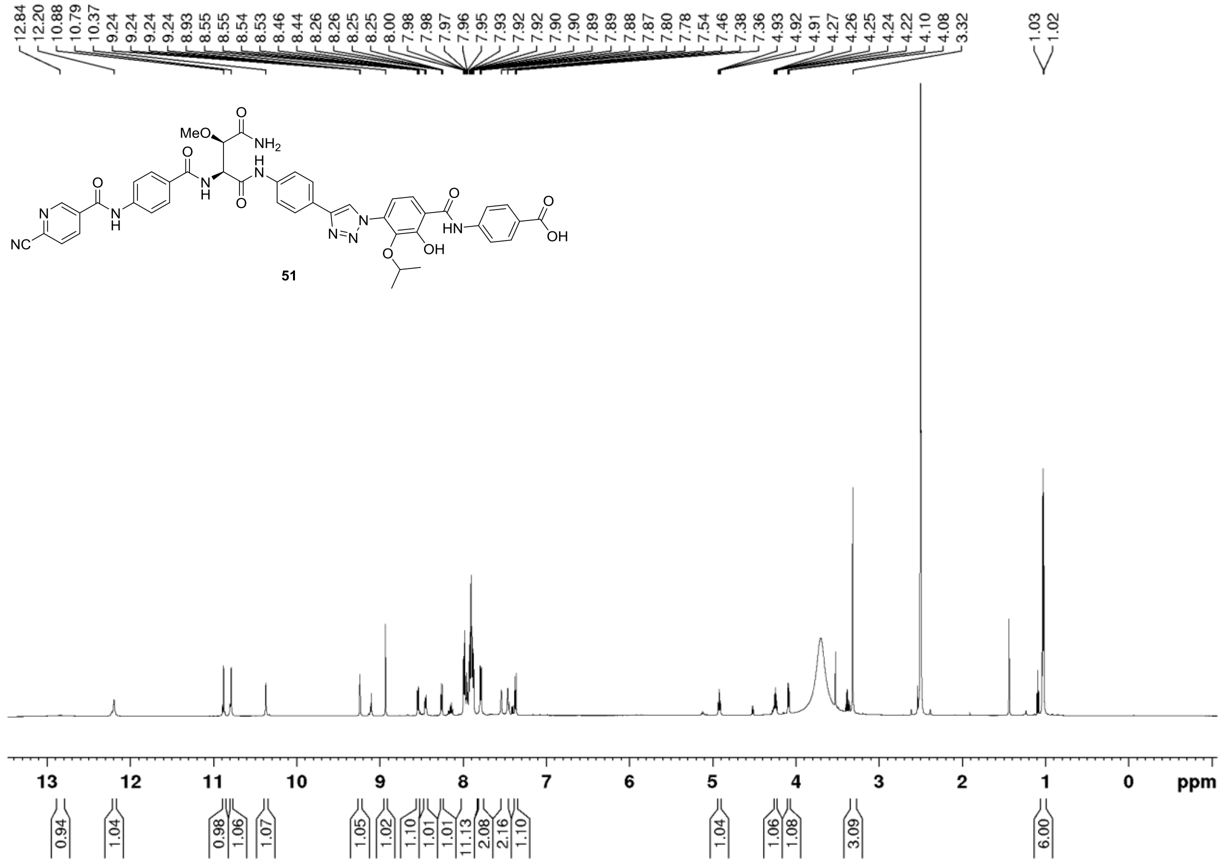
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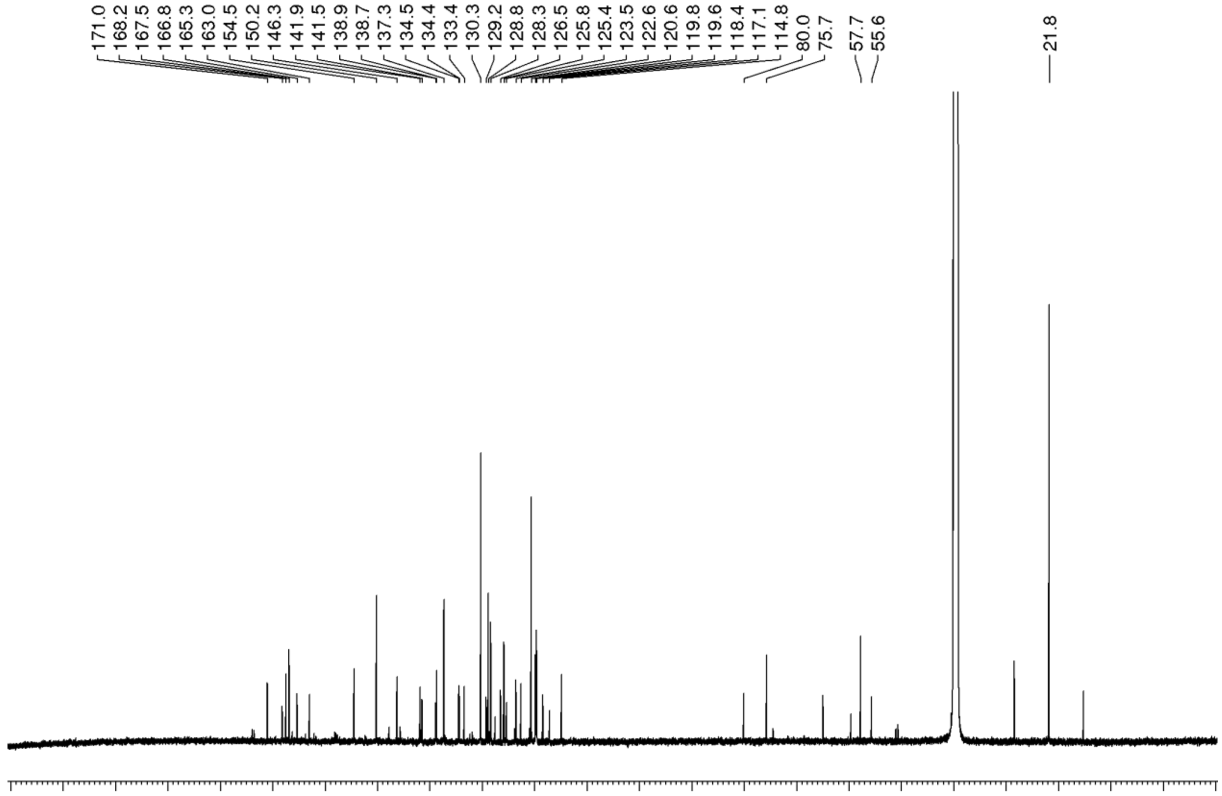
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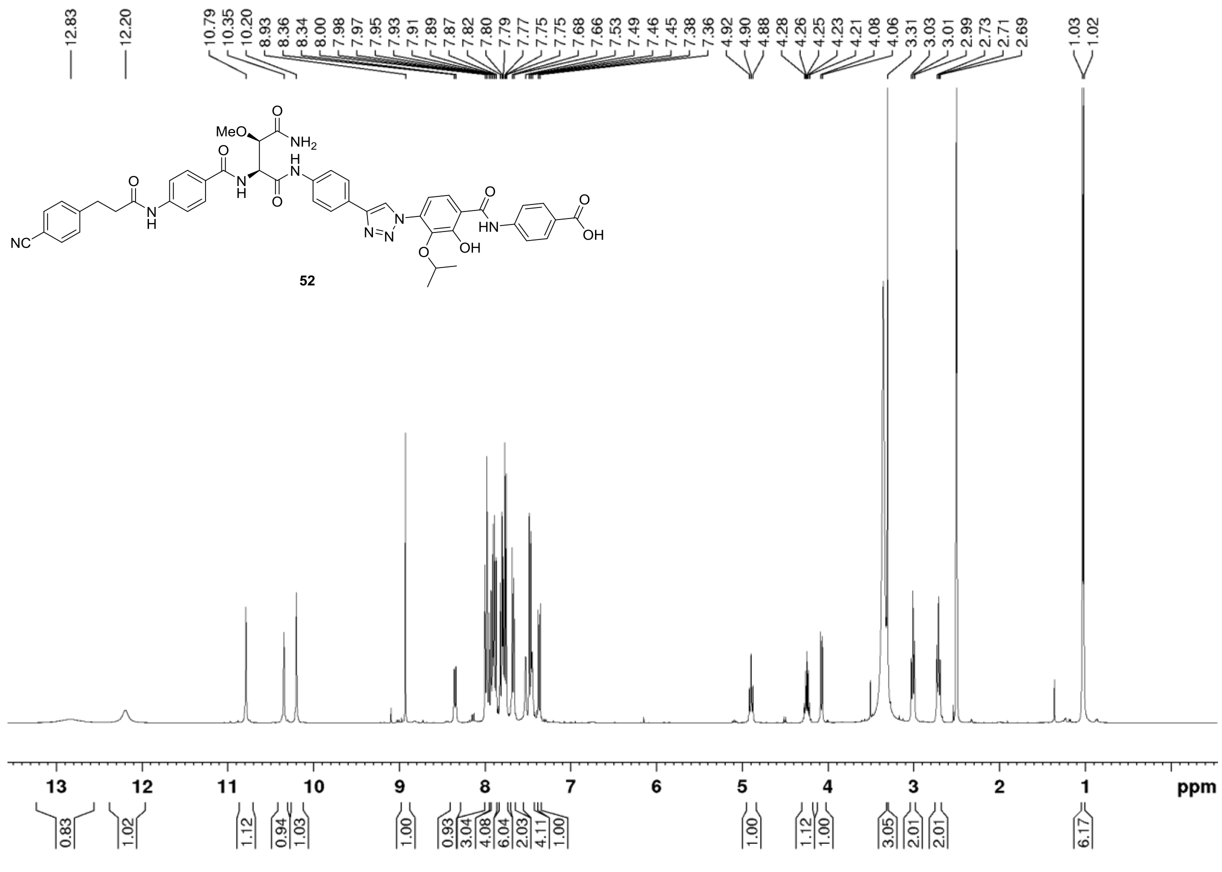
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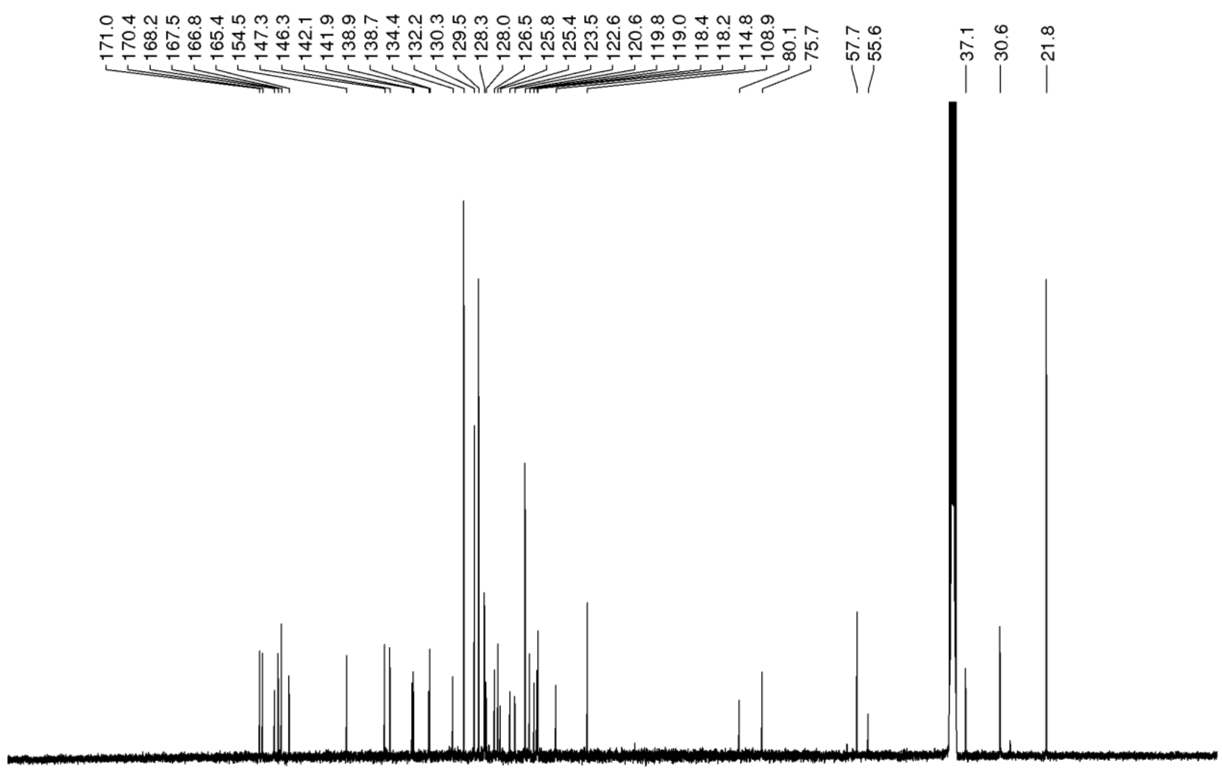
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