

**Modulation of Estrogen-Related Receptors Subtype Selectivity: Conversion of an ER $\beta/\gamma$**   
**Selective Agonist to ER $\alpha/\beta/\gamma$  Pan Agonists**

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**(E)-N'-Benzylidenebenzohydrazide (10) [1].**

Pale Yellow crystals; yield: 89% (EtOH). Mp 214-215 °C (Lit. mp 210-212 °C) [1]. LC/MS m/z: 225 [M+H<sup>+</sup>].

**(E)-N'-(4-(Dimethylamino)benzylidene)benzohydrazide (11)[2].**

Orange crystals; yield: 84% (EtOH). Mp 173-175 °C (Lit. mp 175-176 °C) [2]. LC/MS m/z: 268 [M+H<sup>+</sup>].

**(E)-N'-(4-Chlorobenzylidene)benzohydrazide (12)[3]**

White crystals; yield: 88% (EtOH). Mp 166-167 °C (Lit. mp 164-165 °C) [3]. LC/MS m/z: 259 [M+H<sup>+</sup>].

**(E)-N'-(2-Hydroxybenzylidene)benzohydrazide (13)[4]**

White crystals; yield: 69% (EtOH). Mp 160-162 °C (Lit. mp 163-164 °C)[4]. LC/MS m/z: 241 [M+H<sup>+</sup>].

**(E)-N'-(4-Hydroxybenzylidene)benzohydrazide (14)[5]**

Pale brown crystals; yield: 75% (EtOH). Mp 263-265 °C (Lit. mp 267-268 °C)[5]. LC/MS m/z: 241 [M+H<sup>+</sup>].

**(E)-N'-(4-Methoxybenzylidene)benzohydrazide (15)[6]**

White crystals; yield: 83% (EtOH). Mp 159-160 °C (Lit. mp 158 °C) [6]. LC/MS m/z: 255 [M+H<sup>+</sup>].

**(E)-N'-(3-Nitrobenzylidene)benzohydrazide (17)[6]**

Yellow crystals; yield: 86% (EtOH). Mp 189-190 °C (Lit. mp 188 °C)[6]. LC/MS m/z: 270 [M+H<sup>+</sup>].

**(E)-N'-(4-Nitrobenzylidene)benzohydrazide (18)[6]**

Yellow crystals; yield: 88% (EtOH). Mp 236-238 °C (Lit. mp 240-242 °C)[6]. LC/MS m/z: 270 [M+H<sup>+</sup>].

**(E)-N'-Benzylidene-2-hydroxybenzohydrazide (19)[7]**

White crystals; yield: 90% (EtOH). Mp 228-229 °C (Lit. mp 231 °C)[7]. LC/MS m/z: 241 [M+H<sup>+</sup>].

**(E)-N'-Benzylidene-4-chlorobenzohydrazide (20)[6]**

White crystals; yield: 87% (EtOH). Mp 226-228 °C (Lit. mp 230 °C)[6]. LC/MS m/z: 259 [M+H<sup>+</sup>].

**(E)-N'-Benzylidene-4-bromobenzohydrazide (21)[8]**

White crystals; yield: 81% (EtOH). Mp 236-238 °C (Lit. mp 235-237 °C)[8]. LC/MS m/z: 303 [M+H<sup>+</sup>].

**(E)-N'-Benzylidene-4-nitrobenzohydrazide (22)[9]**

Pale yellow crystals; yield: 81% (EtOH). Mp 254-256 °C (Lit. mp 259 °C)[9]. LC/MS m/z: 270 [M+H<sup>+</sup>].

**(E)-N'-Benzylidene-2-methylbenzohydrazide (23)**[10]

White crystals; yield: 77% (EtOH). Mp 169-171 °C (Lit. mp 173-174 °C)[10]. LC/MS m/z: 239 [M+H<sup>+</sup>].

**(E)-N'-(4-(Dimethylamino)benzylidene)-2-methylbenzohydrazide (25)**[11]

Yellow crystals; yield: 71% (EtOH). Mp 149-151 °C (Lit. mp 151-152 °C). LC/MS m/z: 282 [M+H<sup>+</sup>].

**(E)-4-Chloro-N'-(4-chlorobenzylidene)benzohydrazide (26)**[10]

White crystals; yield: 82% (EtOH). Mp 219-220 °C (Lit. mp 220-221 °C)[10]. LC/MS m/z: 293 [M+H<sup>+</sup>].

**(E)-2-Chloro-N'-(2-hydroxybenzylidene)benzohydrazide (28)**[12]

Yellow crystals; yield: 69% (EtOH). Mp 142-143 °C (Lit. mp 145 °C).[12] LC/MS m/z: 275 [M+H<sup>+</sup>].

**(E)-N'-(4-Hydroxybenzylidene)-2-methylbenzohydrazide (30)**[13]

Colorless microcrystals; yield: 72% (EtOH). Mp 155-157 °C. LC/MS m/z: 255 [M+H<sup>+</sup>].

**(E)-4-Chloro-N'-(2-hydroxybenzylidene)benzohydrazide (31)**[14]

Yellow crystals; yield: 76% (EtOH). Mp 236-238 °C (Lit. mp 240 °C)[14]. LC/MS m/z: 275 [M+H<sup>+</sup>].

**(E)-4-Bromo-N'-(2-hydroxybenzylidene)benzohydrazide (32)**[15]

Yellow crystals; yield: 79% (EtOH). Mp 249-251 °C. LC/MS m/z: 319 [M+H<sup>+</sup>].

**(E)-N'-Benzylidene-2-phenylacetohydrazide (34)**[16]

Colorless solid; yield: 58% (EtOH). Mp 151-153 °C (Lit. mp 154 °C). LC/MS m/z: 239 [M+H<sup>+</sup>].

**(E)-4-Nitro-N'-(4-nitrobenzylidene)benzohydrazide (36)**[17]

Orange yellow crystals; yield: 63% (EtOH). Mp 160-162 °C. LC/MS m/z: 315 [M+H<sup>+</sup>].

**(E)-4-Nitro-N'-(3-nitrobenzylidene)benzohydrazide (37)**[18]

Orange yellow crystals; yield: 63% (EtOH). Mp 189-190 °C. LC/MS m/z: 315 [M+H<sup>+</sup>].

**(E)-N'-Ethylidenebenzohydrazide (39)**[1]

Yellow crystals; yield: 75% (EtOH). Mp 154-156 °C (Lit. mp 159-162 °C)[1]. LC/MS m/z: 163 [M+H<sup>+</sup>].

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**N'-(1*E*,2*E*)-3-Phenylallylidene)benzohydrazide (40)**[19]

Yellow crystals; yield: 53% (EtOH). Mp 186-189 °C (Lit. mp 192-194 °C)[19]. LC/MS m/z: 251 [M+H<sup>+</sup>].

***N'*-((1*E*,2*E*)-3-(4-(Dimethylamino)phenyl)allylidene)benzohydrazide (41)[20]**

Yellow solid; yield: 49% (EtOH). Mp 186-189 °C (Lit. mp 192-194 °C)[19]. LC/MS m/z: 294 [M+H<sup>+</sup>].

**(E)-*N'*-(Furan-2-ylmethylene)benzohydrazide (43)[21]**

Yellow solid; yield: 83% (EtOH). Mp 129-130 °C (Lit. mp 133.8 °C)[21]. LC/MS m/z: 215 [M+H<sup>+</sup>].

**(E)-*N'*-((5-Methylfuran-2-yl)methylene)benzohydrazide (44)[21]**

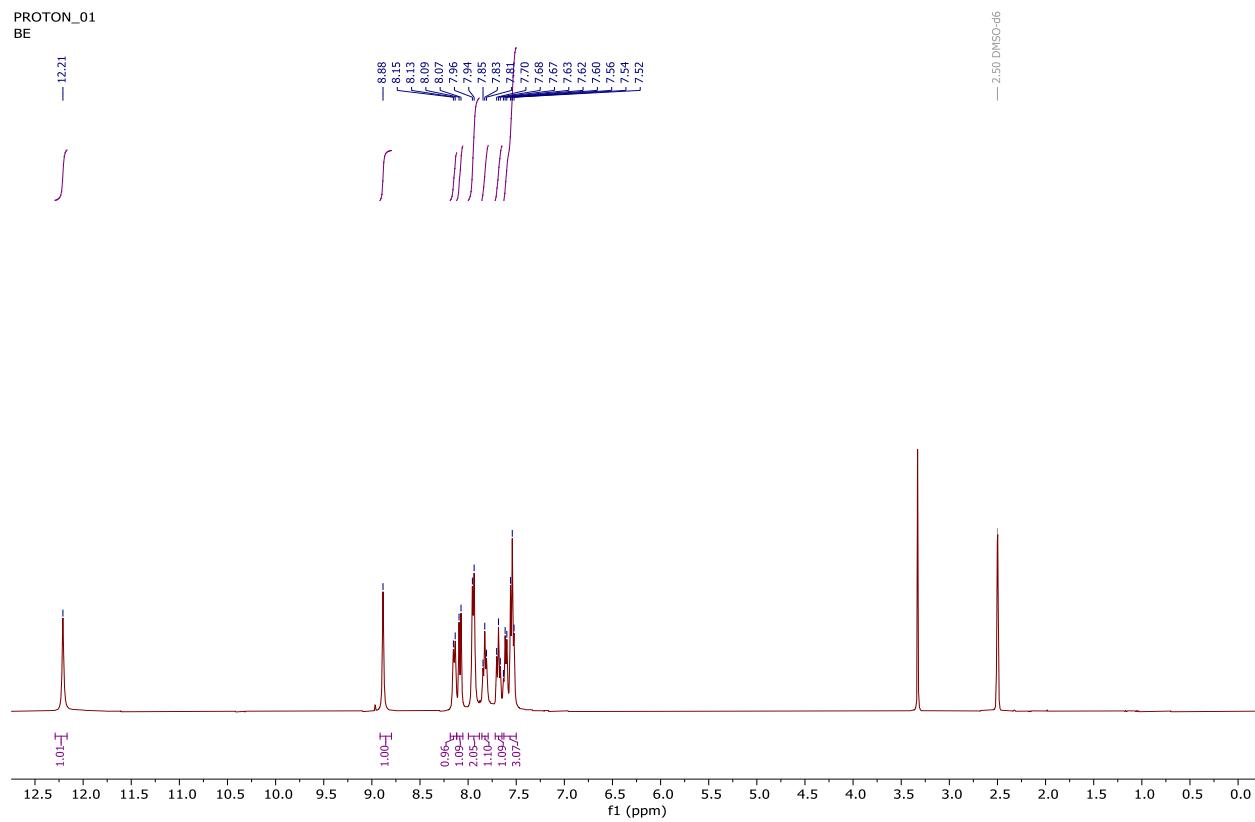
Yellow solid; yield: 85% (EtOH). Mp 122-123 °C. LC/MS m/z: 229 [M+H<sup>+</sup>].

**(E)-*N'*-(Quinolin-2-ylmethylene)benzohydrazide (45)[21]**

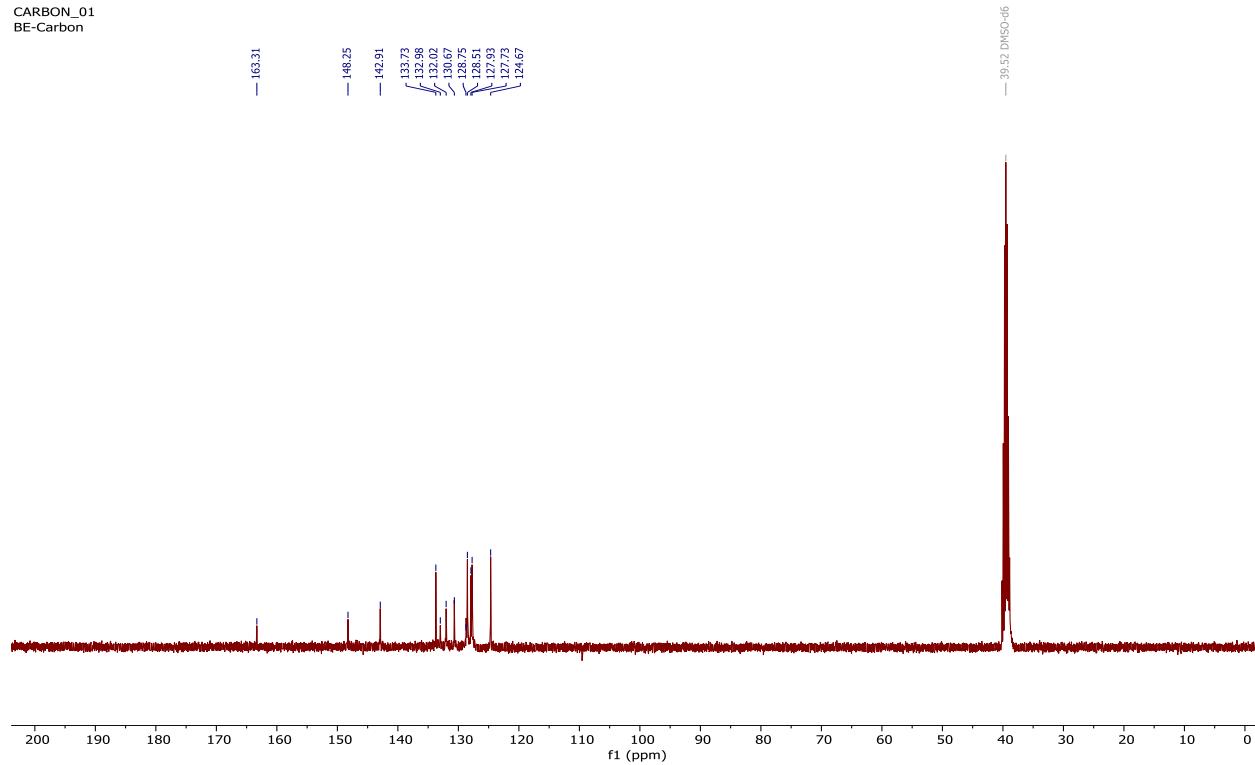
Yellow solid; yield: 80% (EtOH). Mp 172-174 °C (Lit. mp 170 °C)[21]. LC/MS m/z: 276 [M+H<sup>+</sup>].

**(E)-*N'*-(Quinolin-4-ylmethylene)benzohydrazide (46)[21]**

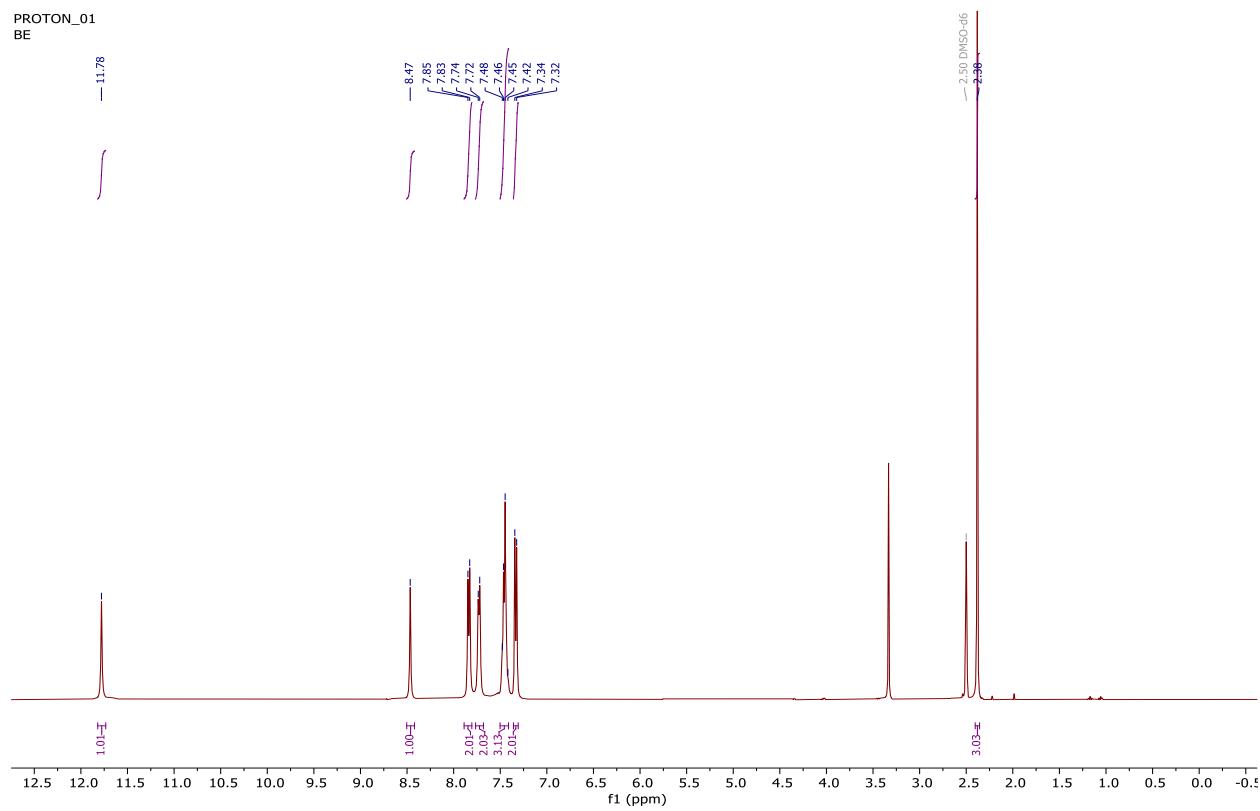
Yellow solid; yield: 84% (EtOH). Mp 214-216 °C (Lit. mp 218 °C)[22]. LC/MS m/z: 276 [M+H<sup>+</sup>].



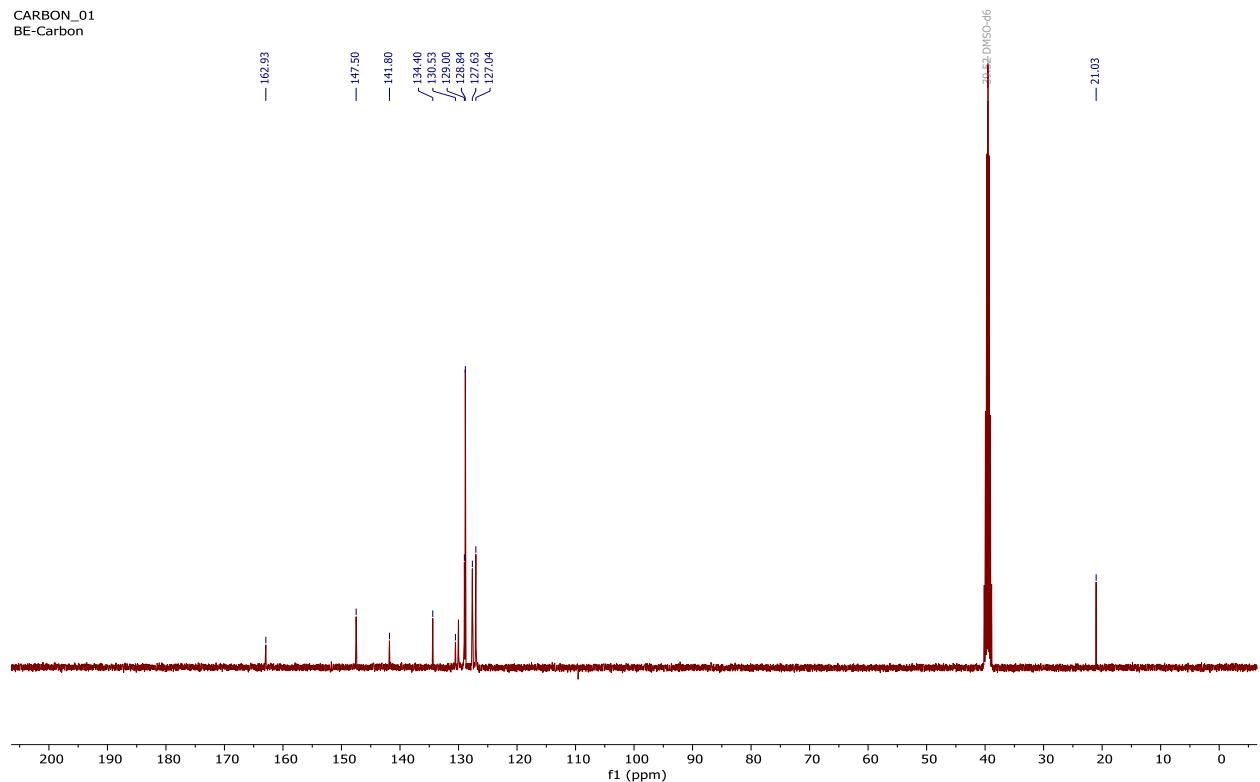
**Figure S1.**  $^1\text{H}$  NMR of compound **16**.



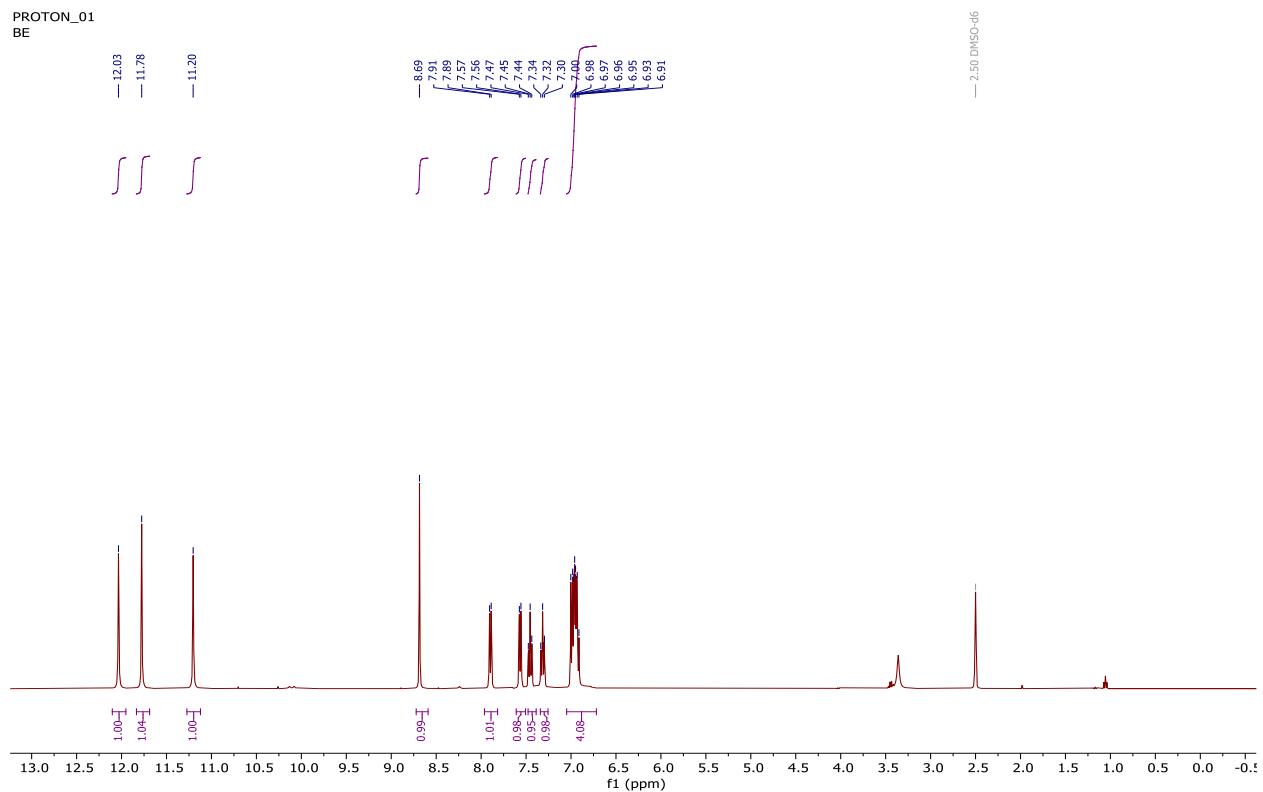
**Figure S2.**  $^{13}\text{C}$  NMR of compound **16**.



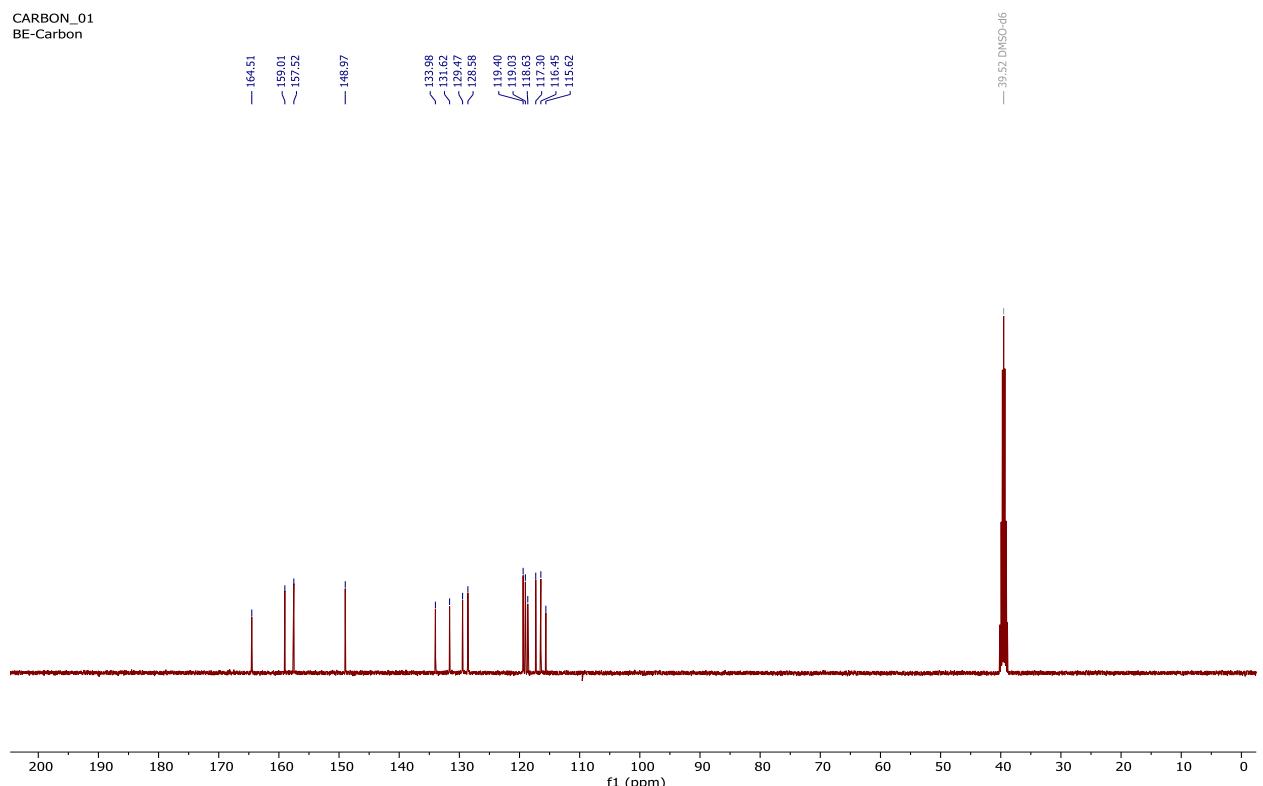
**Figure S3.**  $^1\text{H}$  NMR of compound **24**.



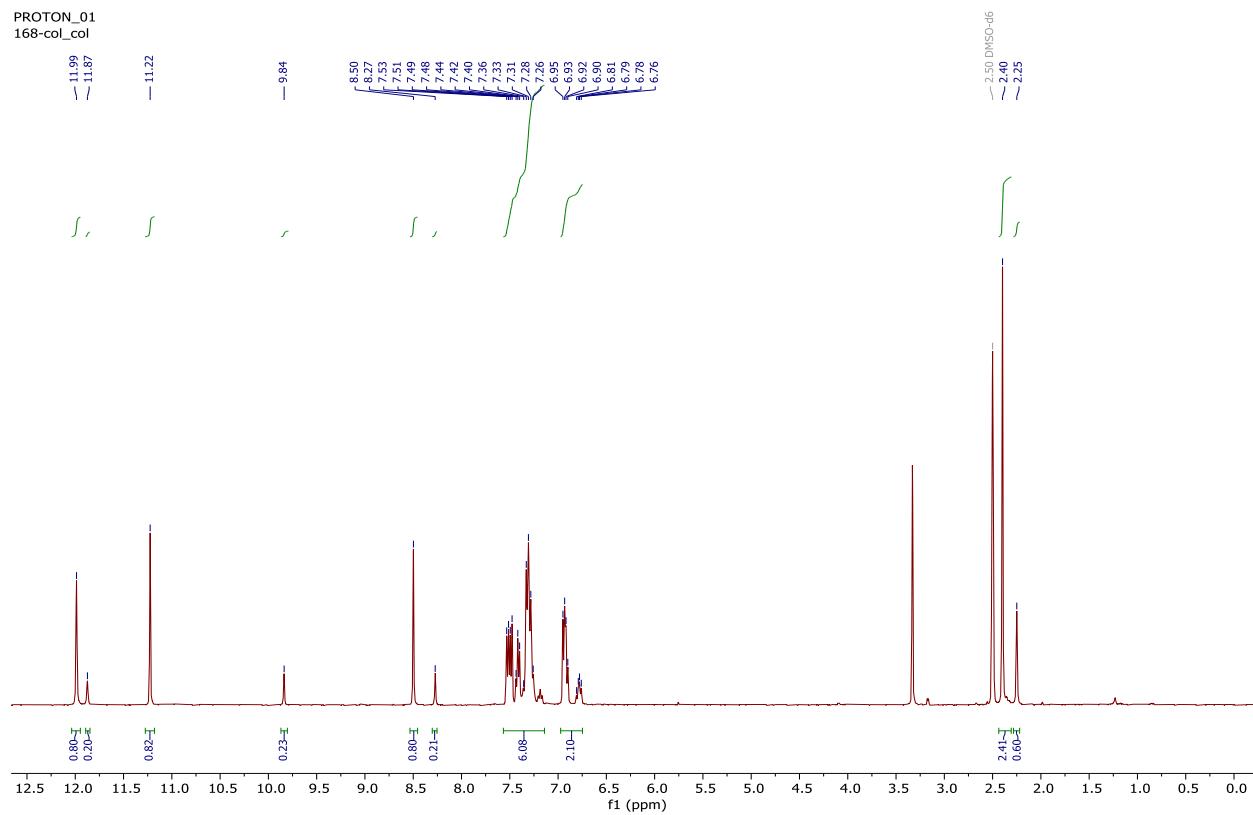
**Figure S4.**  $^{13}\text{C}$  NMR of compound **24**.



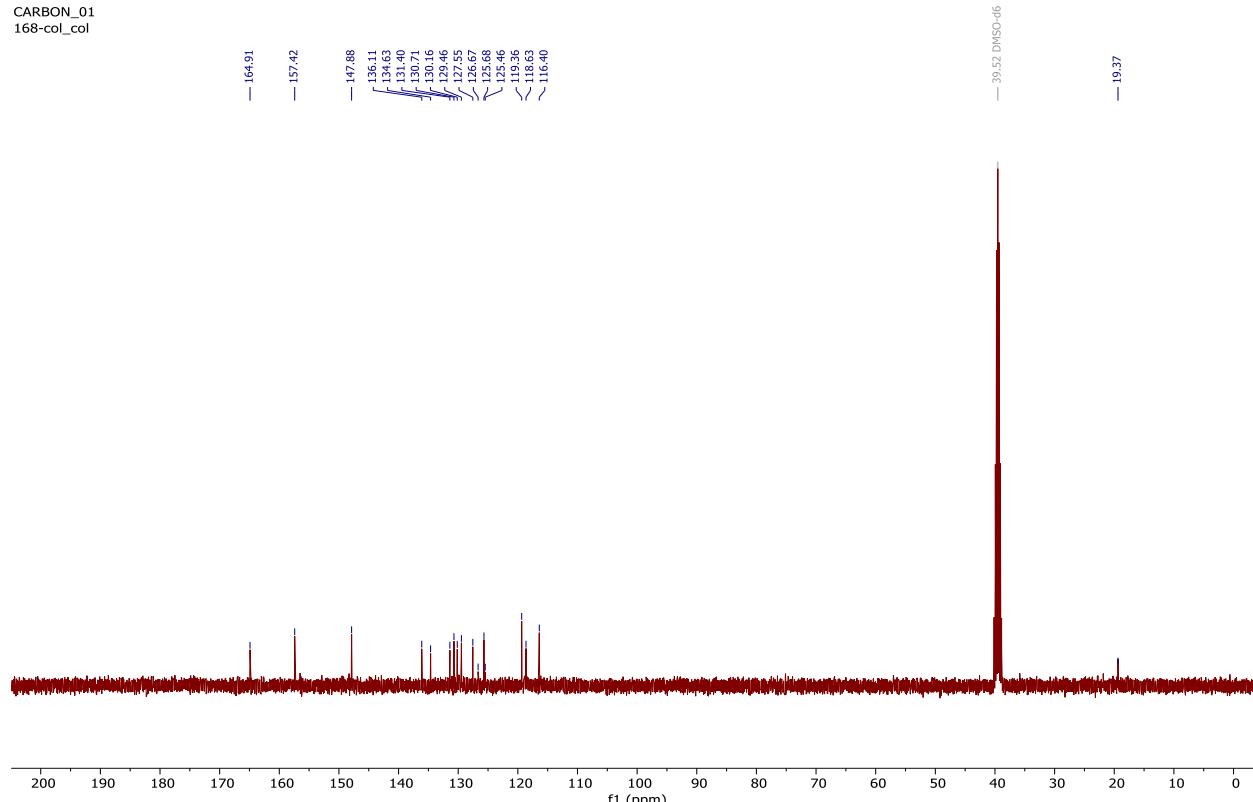
**Figure S5.**  $^1\text{H}$  NMR of compound 27.



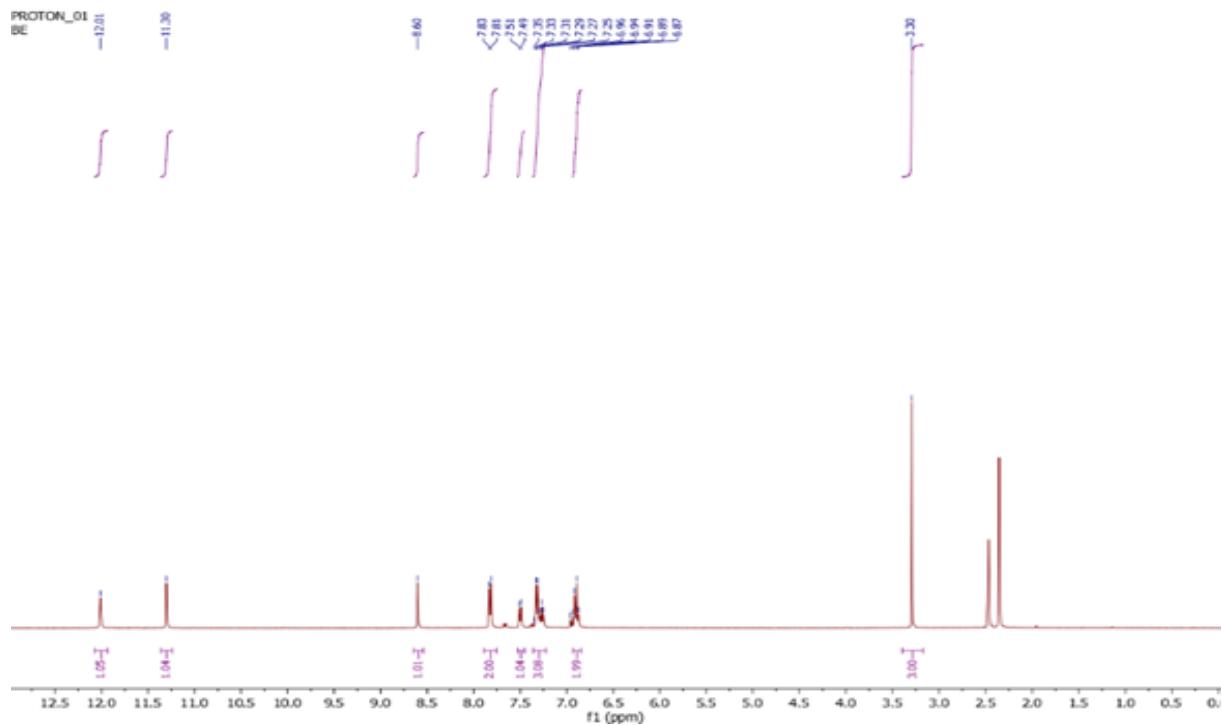
**Figure S6.**  $^{13}\text{C}$  NMR of compound 27.



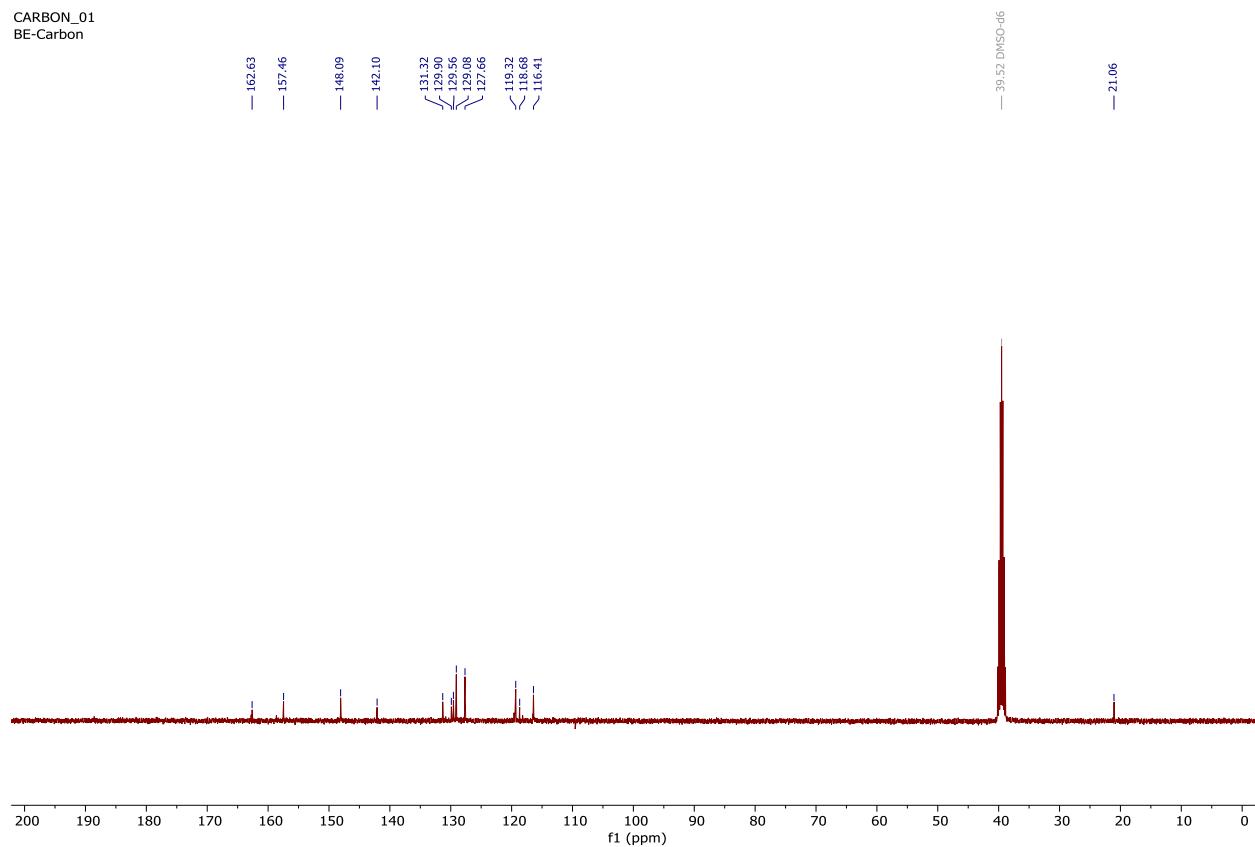
**Figure S7.**  $^1\text{H}$  NMR of compound 29.



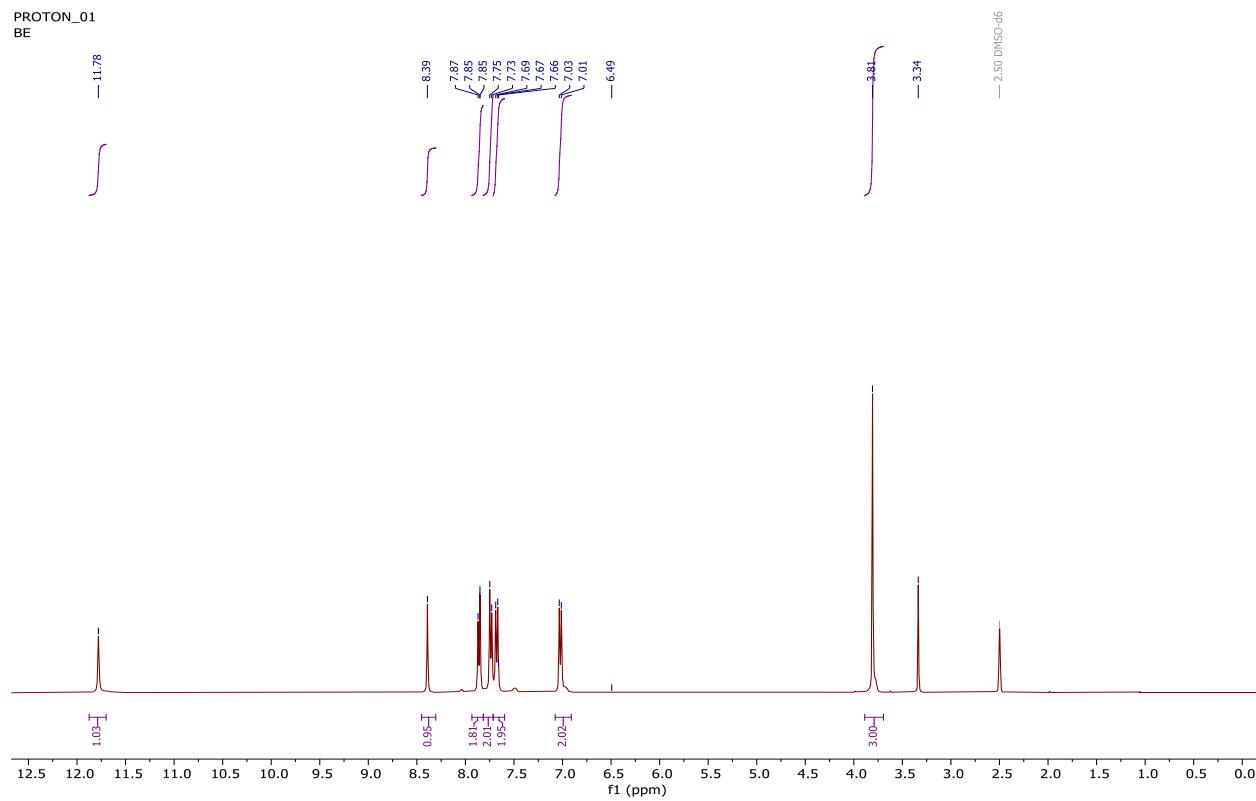
**Figure S8.**  $^{13}\text{C}$  NMR of compound 29.



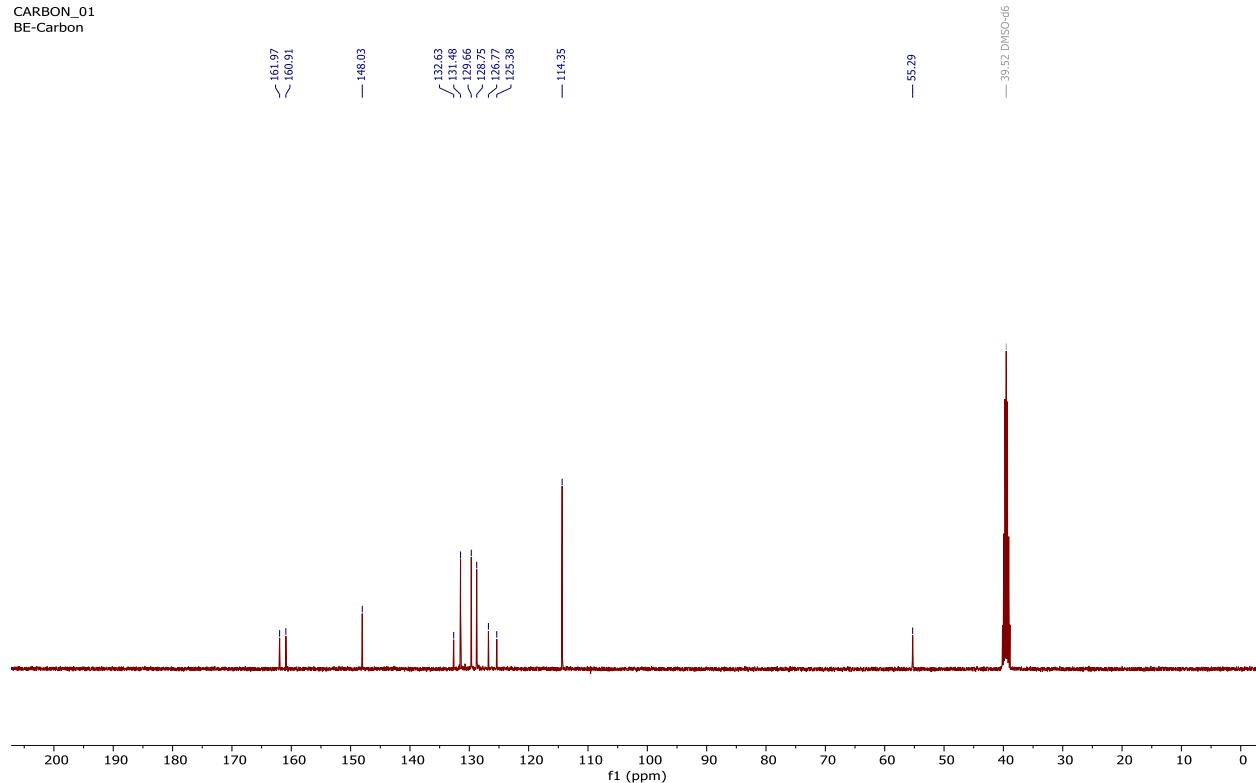
**Figure S9.**  $^1\text{H}$  NMR of compound 33.



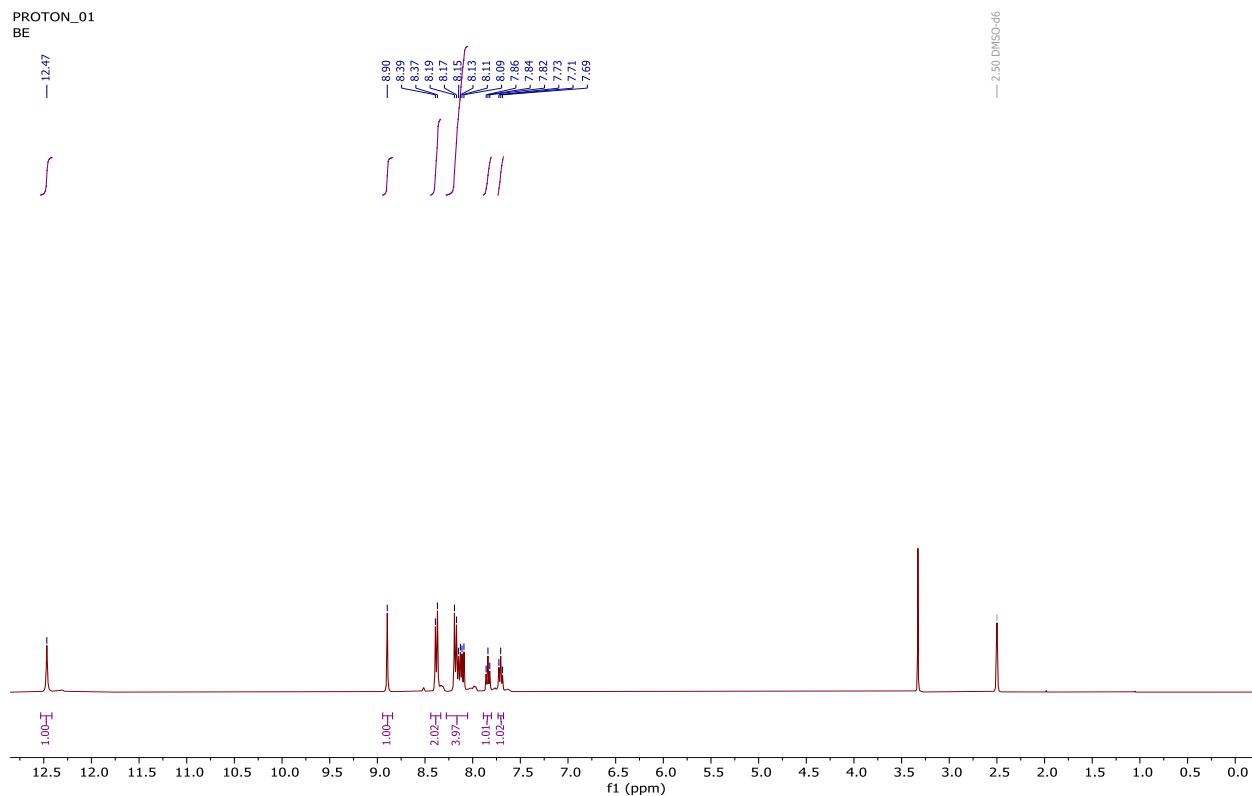
**Figure S10.**  $^{13}\text{C}$  NMR of compound 33.



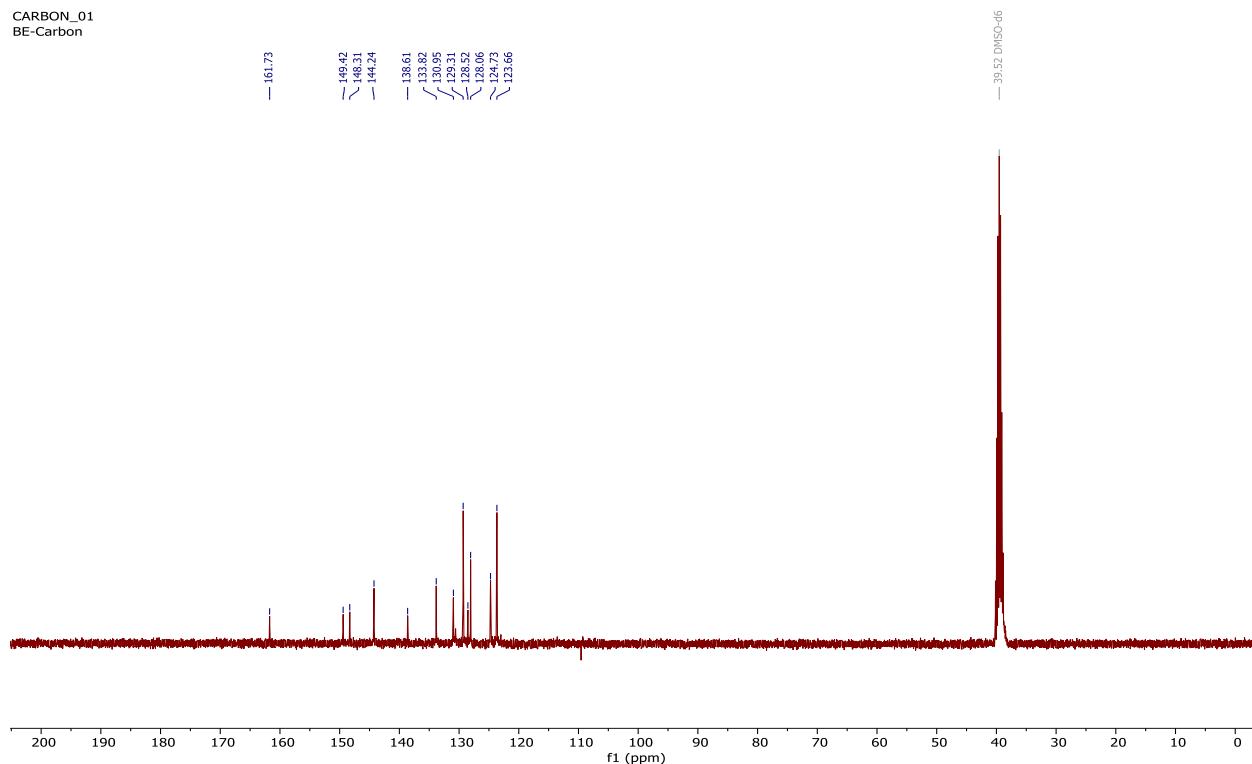
**Figure S11.** <sup>1</sup>H NMR of compound 35.



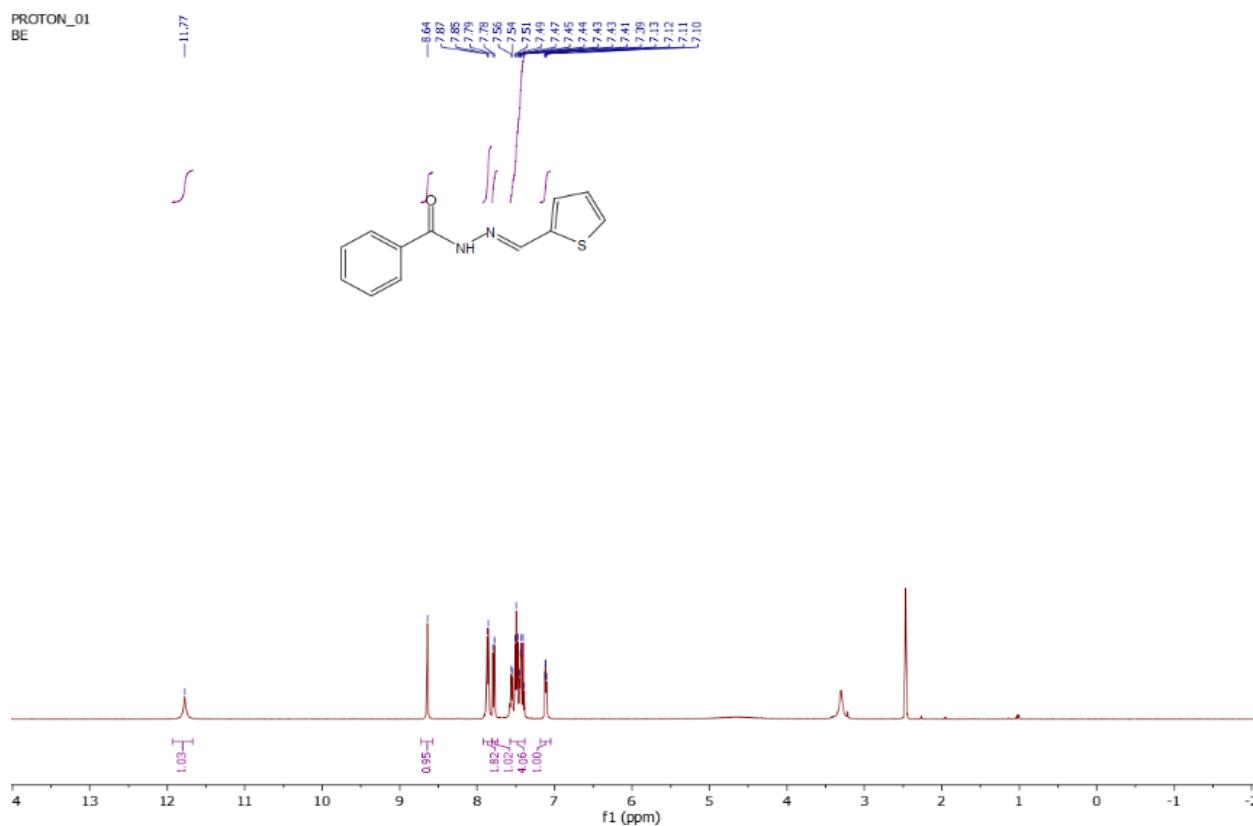
**Figure S12.** <sup>13</sup>C NMR of compound 35.



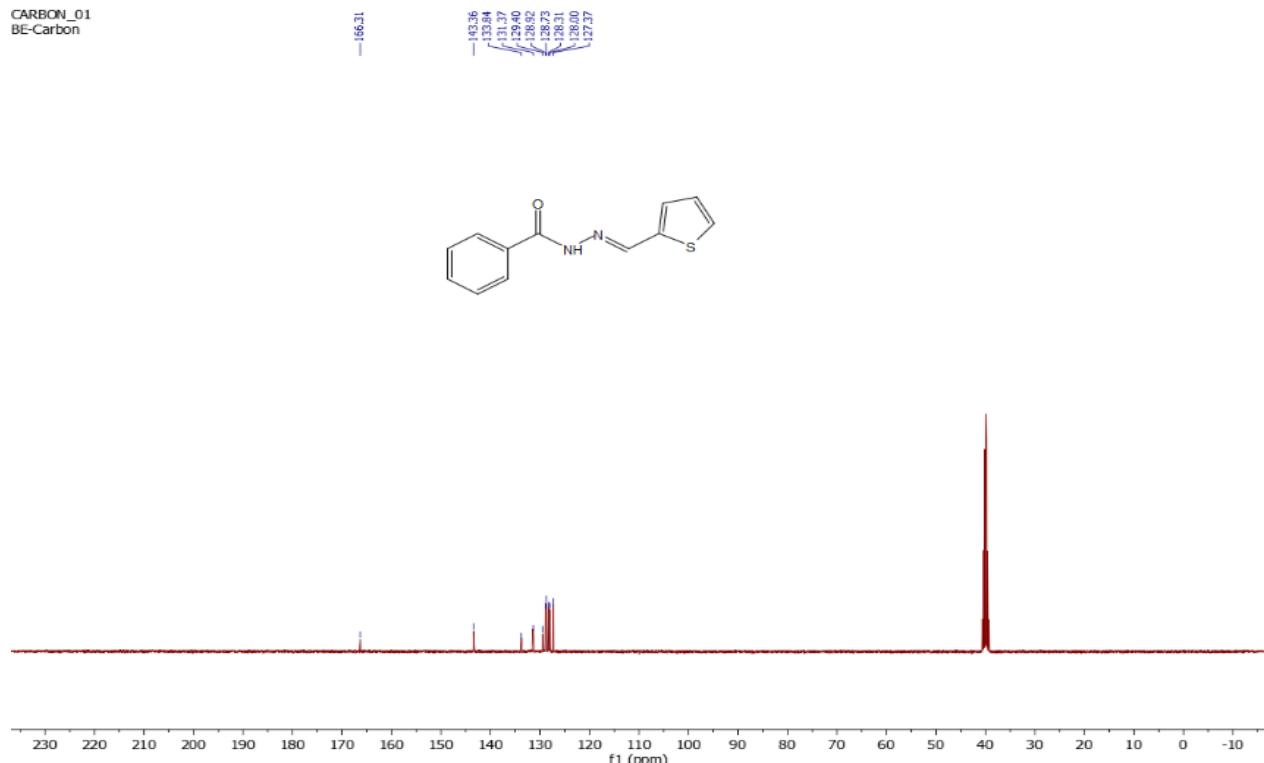
**Figure S13.**  $^1\text{H}$  NMR of compound 38.



**Figure S14.**  $^{13}\text{C}$  NMR of compound 38.



**Figure S15.**  $^1\text{H}$  NMR of compound 42.



**Figure S16.**  $^{13}\text{C}$  NMR of compound 42.

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