

**Rapid incorporation of difluoroacetate radical into *para*-quinone methides via radical 1,6-conjugate addition**

Xin Li,<sup>a</sup> Songtao He,<sup>a</sup> and Qiuling Song <sup>\*a,b</sup>

<sup>a</sup>Institute of Next Generation Matter Transformation, College of Materials Science & Engineering, Huaqiao University, 668 Jimei Blvd, Xiamen 361021, Fujian, China.

<sup>b</sup>State Key Laboratory of Organometallic Chemistry and Key Laboratory of Organofluorine Chemistry, Shanghai Institute of Organic Chemistry, Chinese Academy of Sciences, Shanghai 200032, China.

<sup>\*</sup>qsong@hqu.edu.cn

**Supporting Information**

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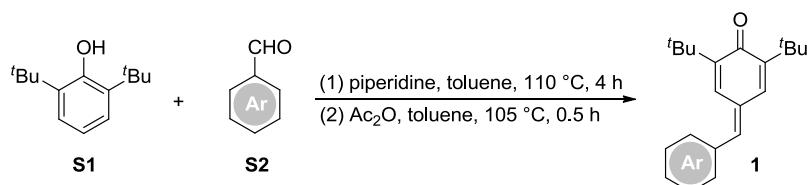
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**I. General Information**

All air or moisture sensitive reactions were conducted in oven-dried glassware under nitrogen atmosphere using dry solvents. Anhydrous THF were freshly distilled from sodium. Unless otherwise stated, chemicals and reagents were used as received. Flash column chromatography was performed over silica gel (200-300 mesh) purchased from *Qindao Bangkai Co., China*. <sup>1</sup>H and <sup>13</sup>C NMR spectra were recorded on a Bruker AV 500 MHz NMR spectrometer using residue solvent peaks as an internal standard (<sup>1</sup>H NMR: CHCl<sub>3</sub> at 7.26 ppm, <sup>13</sup>C NMR: CDCl<sub>3</sub> at 77.0 ppm). HRMS were recorded on an Agilent 6545 Q-TOF LC/MS instrument with electrospray ionization (ESI) technique.

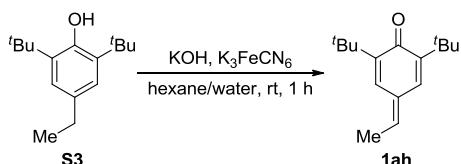
## II. Preparation of Materials

### 2.1. Synthesis of *p*-Quinone Methide Substrates



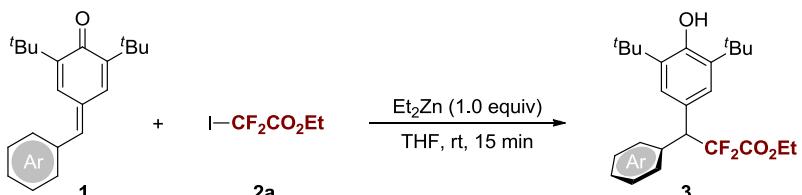
*p*-Quinone methide substrates **1a-1ag**, **1ai**, **1aj** were synthesized according to the literature reported procedure.<sup>1</sup> They are known compounds.

### 2.2. Synthesis of *p*-Quinone Methide **1ah**



*p*-Quinone methide **1ah** was synthesized according to the literature reported procedure, and it is a known compound.<sup>2</sup>

## III. Radical 1,6-Conjugate Addition of *p*-Quinone Methides

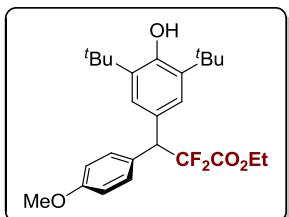


An oven-dried Schlenk tube was charged with *p*-quinone methide substrates (0.1 mmol) and a stir bar. The tube was degassed and refilled with nitrogen, and then a solution of ethyl iododifluoroacetate (0.1 mmol, 1.0 equiv) in anhydrous THF (1 mL) was injected to the tube via a syringe, followed by the addition of a solution of diethylzinc (1.0 mol/L in hexane, 0.1 mL, 1.0 equiv). After stirring at room temperature for 15 minutes, the reaction was quenched with saturated NH<sub>4</sub>Cl and the aqueous layer was extracted with ethyl acetate (3 × 20 mL). The combined organic layers were dried with Na<sub>2</sub>SO<sub>4</sub>, filtered, and concentrated under reduced pressure. The residue was purified by flash silica gel column chromatography (petroleum ether/ethyl acetate) to afford the pure product.

<sup>1</sup> D. Richter, N. Hampel, T. Singer, A. R. Ofial, H. Mayr, *Eur. J. Org. Chem.* **2009**, *19*, 3203.

<sup>2</sup> Z.-Q. Liu, P.-S. You, L.-D. Zhang, D.-Q. Liu, S.-S. Liu, X.-Y. Guan, *Molecules* **2020**, *25*, 539.

**Ethyl 3-(3,5-di-*tert*-butyl-4-hydroxyphenyl)-2,2-difluoro-3-(4-methoxyphenyl)propanoate (3a)**



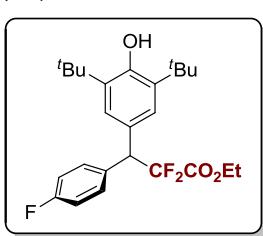
**<sup>1</sup>H NMR** (500 MHz, CDCl<sub>3</sub>) δ 7.37 (d, *J* = 8.7 Hz, 2H), 7.20 (s, 2H), 6.89 (d, *J* = 8.7 Hz, 2H), 5.19 (s, 1H), 4.62 (t, *J* = 18.5 Hz, 1H), 4.14 (qd, *J* = 7.2, 5.3 Hz, 2H), 3.81 (s, 3H), 1.43 (s, 18H), 1.07 (t, *J* = 7.2 Hz, 3H).

**<sup>13</sup>C NMR** (125 MHz, CDCl<sub>3</sub>) δ 164.1 (t, *J* = 32.6 Hz), 159.0, 153.2, 135.7, 130.7, 128.16, 128.14, 126.2, 116.2 (t, *J* = 255.5 Hz), 113.8, 62.5, 55.2, 54.7 (t, *J* = 21.8 Hz), 34.3, 30.2, 13.6.

**<sup>19</sup>F NMR** (471 MHz, CDCl<sub>3</sub>) δ -106.17 (d, *J* = 254.6 Hz), -107.09 (d, *J* = 254.6 Hz).

**HRMS:** (ESI<sup>+</sup>) [M+Na]<sup>+</sup> calcd for C<sub>26</sub>H<sub>34</sub>F<sub>2</sub>O<sub>4</sub>Na: 471.2317, found: 471.2325.

**Ethyl 3-(3,5-di-*tert*-butyl-4-hydroxyphenyl)-2,2-difluoro-3-(4-fluorophenyl)propanoate (3b)**



**<sup>1</sup>H NMR** (500 MHz, CDCl<sub>3</sub>) δ 7.42 (dd, *J* = 8.5, 5.4 Hz, 2H), 7.17 (s, 2H), 7.05 (t, *J* = 8.7 Hz, 1H), 5.22 (s, 1H), 4.66 (t, *J* = 18.2 Hz, 1H), 4.19 – 4.10 (m, 2H), 1.43 (s, 18H), 1.08 (t, *J* = 7.1 Hz, 3H).

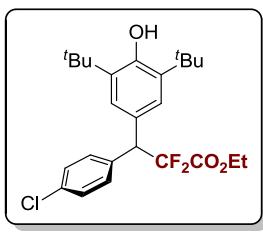
**<sup>13</sup>C NMR** (125 MHz, CDCl<sub>3</sub>) δ 163.9 (t, *J* = 32.5 Hz), 162.2 (d, *J* = 246.3 Hz), 153.4, 135.9, 131.9, 131.2 (d, *J* = 8.1 Hz), 126.1, 125.7 (d, *J* = 4.8 Hz), 116.0 (t, *J* = 256.1 Hz), 115.3 (d, *J* = 21.3 Hz), 62.6, 54.6

(t, *J* = 21.9 Hz), 34.3, 30.2, 13.6.

**<sup>19</sup>F NMR** (471 MHz, CDCl<sub>3</sub>) δ -105.52 (d, *J* = 256.0 Hz), -107.67 (d, *J* = 256.0 Hz), -114.96.

**HRMS:** (ESI<sup>+</sup>) [M+Na]<sup>+</sup> calcd for C<sub>25</sub>H<sub>31</sub>F<sub>3</sub>O<sub>3</sub>Na: 459.2118, found: 459.2117.

**Ethyl 3-(4-chlorophenyl)-3-(3,5-di-*tert*-butyl-4-hydroxyphenyl)-2,2-difluoropropanoate (3c)**



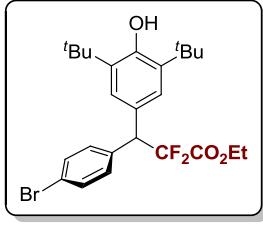
**<sup>1</sup>H NMR** (500 MHz, CDCl<sub>3</sub>) δ 7.38 (d, *J* = 8.6 Hz, 2H), 7.33 (d, *J* = 8.6 Hz, 2H), 7.16 (s, 2H), 5.22 (s, 1H), 4.65 (dd, *J* = 19.1, 17.3 Hz, 1H), 4.23 – 4.08 (m, 2H), 1.43 (s, 18H), 1.08 (t, *J* = 7.1 Hz, 3H).

**<sup>13</sup>C NMR** (125 MHz, CDCl<sub>3</sub>) δ 163.8 (t, *J* = 32.5 Hz), 153.5, 135.9, 134.71, 134.68, 133.6, 130.9, 128.6, 126.1, 125.42, 125.38, 115.9 (t, *J* = 255.3 Hz), 62.7, 54.8 (t, *J* = 21.9 Hz), 34.3, 30.2, 13.6.

**<sup>19</sup>F NMR** (471 MHz, CDCl<sub>3</sub>) δ -105.06 (d, *J* = 256.4 Hz), -107.88 (d, *J* = 256.4 Hz).

**HRMS:** (ESI<sup>+</sup>) [M+Na]<sup>+</sup> calcd for C<sub>25</sub>H<sub>31</sub>ClF<sub>2</sub>O<sub>3</sub>Na: 475.1822, found: 475.1826.

**Ethyl 3-(4-bromophenyl)-3-(3,5-di-*tert*-butyl-4-hydroxyphenyl)-2,2-difluoropropanoate (3d)**



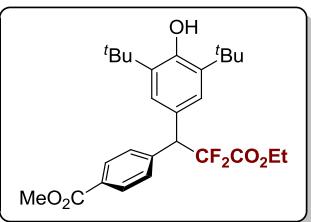
**<sup>1</sup>H NMR** (500 MHz, CDCl<sub>3</sub>) δ 7.48 (d, *J* = 8.4 Hz, 2H), 7.32 (d, *J* = 8.4 Hz, 2H), 7.16 (s, 2H), 5.22 (s, 1H), 4.64 (dd, *J* = 19.2, 17.1 Hz, 1H), 4.21 – 4.09 (m, 2H), 1.43 (s, 18H), 1.08 (t, *J* = 7.1 Hz, 3H).

**<sup>13</sup>C NMR** (125 MHz, CDCl<sub>3</sub>) δ 163.8 (t, *J* = 32.4 Hz), 153.5, 136.0, 135.26, 135.23, 131.6, 131.3, 126.1, 125.4, 125.3, 121.75, 115.8 (t, *J* = 256.4 Hz), 62.7, 54.8 (t, *J* = 22.0 Hz), 34.3, 30.2, 13.6.

**<sup>19</sup>F NMR** (471 MHz, CDCl<sub>3</sub>) δ -104.90 (d, *J* = 256.5 Hz), -107.98 (d, *J* = 256.5 Hz).

**HRMS:** (ESI<sup>+</sup>) [M+Na]<sup>+</sup> calcd for C<sub>25</sub>H<sub>31</sub>BrF<sub>2</sub>O<sub>3</sub>Na: 475.1822, found: 475.1826.

**Methyl 4-(1-(3,5-di-*tert*-butyl-4-hydroxyphenyl)-3-ethoxy-2,2-difluoro-3-oxopropyl)benzoate (3e)**



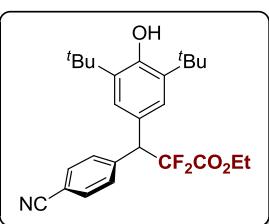
**<sup>1</sup>H NMR** (500 MHz, CDCl<sub>3</sub>) δ 8.03 (d, *J* = 8.2 Hz, 2H), 7.53 (d, *J* = 8.2 Hz, 2H), 7.16 (s, 2H), 5.23 (s, 1H), 4.74 (t, *J* = 18.1 Hz, 1H), 4.14 (qt, *J* = 7.1, 3.6 Hz, 2H), 3.93 (s, 3H), 1.42 (s, 18H), 1.07 (t, *J* = 7.1 Hz, 3H).

**<sup>13</sup>C NMR** (125 MHz, CDCl<sub>3</sub>) δ 166.8, 163.8 (t, *J* = 32.8 Hz), 153.5, 141.34, 141.31, 135.9, 129.7, 129.6, 129.3, 126.2, 125.13, 125.09, 115.8 (t, *J* = 255.9 Hz), 62.7, 55.3 (t, *J* = 21.8 Hz), 52.1, 34.3, 30.2, 13.6.

**<sup>19</sup>F NMR** (471 MHz, CDCl<sub>3</sub>) δ -105.14 (d, *J* = 256.9 Hz), -107.39 (d, *J* = 256.9 Hz).

**HRMS:** (ESI<sup>+</sup>) [M+Na]<sup>+</sup> calcd for C<sub>27</sub>H<sub>34</sub>F<sub>2</sub>O<sub>5</sub>Na: 499.2267, found: 499.2270.

### Ethyl 3-(4-cyanophenyl)-3-(3,5-di-tert-butyl-4-hydroxyphenyl)-2,2-difluoropropanoate (3f)



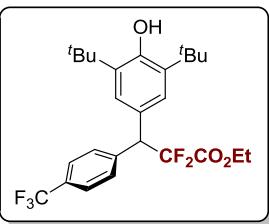
**<sup>1</sup>H NMR** (500 MHz, CDCl<sub>3</sub>) δ 7.65 (d, *J* = 8.2 Hz, 2H), 7.55 (d, *J* = 8.2 Hz, 2H), 7.13 (s, 2H), 5.26 (s, 1H), 4.73 (dd, *J* = 20.6, 15.0 Hz, 1H), 4.22 – 4.09 (m, 2H), 1.42 (s, 18H), 1.08 (t, *J* = 7.1 Hz, 3H).

**<sup>13</sup>C NMR** (125 MHz, CDCl<sub>3</sub>) δ 163.5 (t, *J* = 32.3 Hz), 153.7, 141.63, 141.60, 136.2, 132.2, 130.4, 126.1, 124.61, 124.56, 118.6, 115.6 (dd, *J* = 255.4, 255.0 Hz), 111.6, 62.8, 55.3 (t, *J* = 21.9 Hz), 34.4, 30.2, 13.6.

**<sup>19</sup>F NMR** (471 MHz, CDCl<sub>3</sub>) δ -103.22 (d, *J* = 258.2 Hz), -109.00 (d, *J* = 258.2 Hz).

**HRMS:** (ESI<sup>+</sup>) [M+Na]<sup>+</sup> calcd for C<sub>26</sub>H<sub>31</sub>F<sub>2</sub>NO<sub>3</sub>Na: 466.2164, found: 466.2165.

### Ethyl 3-(3,5-di-tert-butyl-4-hydroxyphenyl)-2,2-difluoro-3-(4-(nitrophenoxy)phenyl)propanoate (3g)



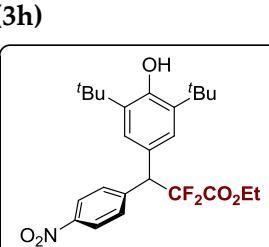
**<sup>1</sup>H NMR** (500 MHz, CDCl<sub>3</sub>) δ 7.62 (d, *J* = 8.3 Hz, 2H), 7.58 (d, *J* = 8.3 Hz, 2H), 7.18 (s, 2H), 5.25 (s, 1H), 4.74 (dd, *J* = 20.2, 15.9 Hz, 1H), 4.26 – 4.06 (m, 2H), 1.43 (s, 18H), 1.07 (t, *J* = 7.1 Hz, 3H).

**<sup>13</sup>C NMR** (125 MHz, CDCl<sub>3</sub>) δ 163.7 (t, *J* = 32.3 Hz), 153.6, 140.3, 140.2, 136.1, 129.9, 129.8 (q, *J* = 32.7 Hz), 126.2, 125.4 (q, *J* = 3.8 Hz), 125.04, 125.00, 124.0 (q, *J* = 272.2 Hz), 115.8 (t, *J* = 255.9 Hz), 62.7, 55.2 (t, *J* = 21.9 Hz), 34.4, 30.2, 13.6.

**<sup>19</sup>F NMR** (471 MHz, CDCl<sub>3</sub>) δ -62.64, -103.99 (d, *J* = 257.2 Hz), -108.56 (d, *J* = 257.2 Hz).

**HRMS:** (ESI<sup>+</sup>) [M+Na]<sup>+</sup> calcd for C<sub>26</sub>H<sub>31</sub>F<sub>5</sub>O<sub>3</sub>Na: 509.2086, found: 509.2093.

### Ethyl 3-(3,5-di-tert-butyl-4-hydroxyphenyl)-2,2-difluoro-3-(4-nitrophenyl)propanoate (3h)



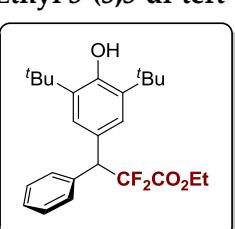
**<sup>1</sup>H NMR** (500 MHz, CDCl<sub>3</sub>) δ 8.22 (d, *J* = 8.8 Hz, 2H), 7.62 (d, *J* = 8.8 Hz, 2H), 7.14 (s, 2H), 5.27 (s, 1H), 4.80 (dd, *J* = 20.9, 14.6 Hz, 1H), 4.25 – 4.10 (m, 2H), 1.43 (s, 18H), 1.09 (t, *J* = 7.1 Hz, 3H).

**<sup>13</sup>C NMR** (125 MHz, CDCl<sub>3</sub>) δ 163.4 (t, *J* = 32.2 Hz), 153.8, 147.3, 143.64, 143.61, 136.3, 130.5, 126.1, 124.54, 124.49, 123.6, 115.6 (t, *J* = 255.1 Hz), 62.9, 55.1 (t, *J* = 21.9 Hz), 34.4, 30.2, 13.6.

**<sup>19</sup>F NMR** (471 MHz, CDCl<sub>3</sub>) δ -102.93 (d, *J* = 258.3 Hz), -109.18 (d, *J* = 258.3 Hz).

**HRMS:** (ESI<sup>+</sup>) [M+Na]<sup>+</sup> calcd for C<sub>25</sub>H<sub>31</sub>F<sub>2</sub>NO<sub>5</sub>Na: 486.2063, found: 486.2064.

### Ethyl 3-(3,5-di-tert-butyl-4-hydroxyphenyl)-2,2-difluoro-3-phenylpropanoate (3i)



**<sup>1</sup>H NMR** (500 MHz, CDCl<sub>3</sub>) δ 7.46 (d, *J* = 7.5 Hz, 2H), 7.38 – 7.33 (m, 2H), 7.33 – 7.29 (m, 1H), 7.22 (s, 2H), 5.20 (s, 1H), 4.67 (t, *J* = 18.5 Hz,

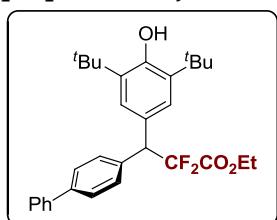
1H), 4.24 – 4.07 (m, 2H), 1.44 (s, 18H), 1.06 (t,  $J$  = 7.1 Hz, 3H).

**$^{13}\text{C}$  NMR** (125 MHz,  $\text{CDCl}_3$ )  $\delta$  164.1 (t,  $J$  = 32.9 Hz), 153.3, 136.21, 136.18, 135.8, 129.6, 128.5, 127.5, 126.3, 126.0, 125.9, 116.2 (t,  $J$  = 255.9 Hz), 62.5, 55.5 (t,  $J$  = 21.8 Hz), 34.3, 30.3, 13.6.

**$^{19}\text{F}$  NMR** (471 MHz,  $\text{CDCl}_3$ )  $\delta$  -105.80 (d,  $J$  = 255.4 Hz), -107.03 (d,  $J$  = 255.4 Hz).

**HRMS:** (ESI $^+$ ) [M+H] $^+$  calcd for  $\text{C}_{25}\text{H}_{33}\text{F}_2\text{O}_3$ : 419.2389, found: 419.2392.

**Ethyl 3-([1,1'-biphenyl]-4-yl)-3-(3,5-di-tert-butyl-4-hydroxyphenyl)-2,2-difluoropropanoate (3j)**



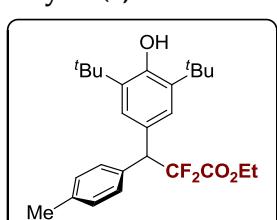
**$^1\text{H}$  NMR** (500 MHz,  $\text{CDCl}_3$ )  $\delta$  7.65 – 7.57 (m, 4H), 7.53 (d,  $J$  = 8.2 Hz, 2H), 7.45 (dd,  $J$  = 8.2, 7.0 Hz, 2H), 7.40 – 7.34 (m, 1H), 7.28 (s, 2H), 5.21 (s, 1H), 4.71 (t,  $J$  = 18.4 Hz, 1H), 4.24 – 4.10 (m, 2H), 1.45 (s, 18H), 1.08 (t,  $J$  = 7.1 Hz, 3H).

**$^{13}\text{C}$  NMR** (125 MHz,  $\text{CDCl}_3$ )  $\delta$  164.1 (t,  $J$  = 32.4 Hz), 153.4, 140.6, 140.4, 135.9, 135.23, 135.20, 130.0, 128.8, 127.3, 127.2, 127.0, 126.3, 125.89, 125.86, 116.2 (t,  $J$  = 255.7 Hz), 62.6, 55.2 (t,  $J$  = 21.8 Hz), 34.4, 30.3, 13.6.

**$^{19}\text{F}$  NMR** (471 MHz,  $\text{CDCl}_3$ )  $\delta$  -105.43 (d,  $J$  = 255.3 Hz), -107.36 (d,  $J$  = 255.3 Hz).

**HRMS:** (ESI $^+$ ) [M+Na] $^+$  calcd for  $\text{C}_{31}\text{H}_{36}\text{F}_2\text{O}_3\text{Na}$ : 495.2705, found: 495.2713.

**Ethyl 3-(3,5-di-tert-butyl-4-hydroxyphenyl)-2,2-difluoropropanoate (3k)**



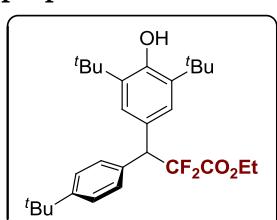
**$^1\text{H}$  NMR** (500 MHz,  $\text{CDCl}_3$ )  $\delta$  7.34 (d,  $J$  = 7.9 Hz, 2H), 7.21 (s, 2H), 7.16 (d,  $J$  = 7.9 Hz, 2H), 5.18 (s, 1H), 4.62 (t,  $J$  = 18.5 Hz, 1H), 4.19 – 4.06 (m, 2H), 2.34 (s, 3H), 1.43 (s, 18H), 1.07 (t,  $J$  = 7.1 Hz, 3H).

**$^{13}\text{C}$  NMR** (125 MHz,  $\text{CDCl}_3$ )  $\delta$  164.1 (t,  $J$  = 32.6 Hz), 153.3, 137.2, 135.7, 133.11, 133.08, 129.4, 129.2, 126.2, 126.12, 126.09, 116.2 (t,  $J$  = 255.6 Hz), 62.5, 55.1 (t,  $J$  = 21.8 Hz), 34.3, 30.2, 21.0, 13.6.

**$^{19}\text{F}$  NMR** (471 MHz,  $\text{CDCl}_3$ )  $\delta$  -105.76 (d,  $J$  = 254.6 Hz), -107.23 (d,  $J$  = 254.6 Hz).

**HRMS:** (ESI $^+$ ) [M+Na] $^+$  calcd for  $\text{C}_{26}\text{H}_{34}\text{F}_2\text{O}_3\text{Na}$ : 455.2368, found: 455.2370.

**Ethyl 3-(4-(tert-butyl)phenyl)-3-(3,5-di-tert-butyl-4-hydroxyphenyl)-2,2-difluoropropanoate (3l)**



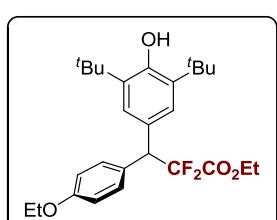
**$^1\text{H}$  NMR** (500 MHz,  $\text{CDCl}_3$ )  $\delta$  7.43 – 7.33 (m, 4H), 7.23 (s, 2H), 5.18 (s, 1H), 4.61 (t,  $J$  = 18.6 Hz, 1H), 4.18 – 4.06 (m, 2H), 1.43 (s, 18H), 1.31 (s, 9H), 1.02 (t,  $J$  = 7.1 Hz, 3H).

**$^{13}\text{C}$  NMR** (125 MHz,  $\text{CDCl}_3$ )  $\delta$  164.2 (t,  $J$  = 32.3 Hz), 153.3, 150.4, 135.7, 133.02, 132.99, 129.2, 126.3, 126.13, 126.09, 125.4, 116.3 (t,  $J$  = 255.8 Hz), 62.4, 55.2 (t,  $J$  = 21.8 Hz), 34.4, 34.3, 31.3, 30.3, 13.6.

**$^{19}\text{F}$  NMR** (471 MHz,  $\text{CDCl}_3$ )  $\delta$  -105.56 (d,  $J$  = 253.5 Hz), -107.76 (d,  $J$  = 253.5 Hz).

**HRMS:** (ESI $^+$ ) [M+Na] $^+$  calcd for  $\text{C}_{29}\text{H}_{40}\text{F}_2\text{O}_3\text{Na}$ : 497.2838, found: 497.2846.

**Ethyl 3-(3,5-di-tert-butyl-4-hydroxyphenyl)-3-(4-ethoxyphenyl)-2,2-difluoropropanoate (3m)**



**$^1\text{H}$  NMR** (500 MHz,  $\text{CDCl}_3$ )  $\delta$  7.35 (d,  $J$  = 8.7 Hz, 2H), 7.19 (s, 2H), 6.87 (d,  $J$  = 8.7 Hz, 2H), 5.18 (s, 1H), 4.60 (t,  $J$  = 18.5 Hz, 1H), 4.13 (qd,  $J$  = 7.2, 4.6 Hz, 2H), 4.03 (q,  $J$  = 7.0 Hz, 2H), 1.43 (s, 18H), 1.42 (t,  $J$  = 7.0 Hz, 3H), 1.07 (t,  $J$  = 7.1 Hz, 3H).

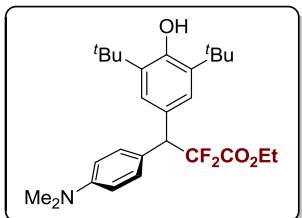
**$^{13}\text{C}$  NMR** (125 MHz,  $\text{CDCl}_3$ )  $\delta$  164.2 (t,  $J$  = 32.8 Hz), 158.4, 153.3, 135.7, 130.7, 127.99, 126.2, 116.2 (t,  $J$  = 256.4 Hz), 114.4, 63.4, 62.5,

54.7 (t,  $J = 22.0$  Hz), 34.3, 30.3, 14.8, 13.7.

$^{19}\text{F}$  NMR (471 MHz,  $\text{CDCl}_3$ )  $\delta$  -106.36 (d,  $J = 254.3$  Hz), -106.94 (d,  $J = 254.3$  Hz).

HRMS: (ESI $^+$ ) [M+Na] $^+$  calcd for  $\text{C}_{27}\text{H}_{36}\text{F}_2\text{O}_4\text{Na}$ : 485.2474, found: 485.2479.

**Ethyl 3-(3,5-di-*tert*-butyl-4-hydroxyphenyl)-3-(4-(dimethylamino)phenyl)-2,2-difluoropropanoate (3n)**



$^1\text{H}$  NMR (500 MHz,  $\text{CDCl}_3$ )  $\delta$  7.31 (d,  $J = 8.7$  Hz, 2H), 7.23 (s, 2H), 6.71 (d,  $J = 8.7$  Hz, 2H), 5.16 (s, 1H), 4.56 (t,  $J = 18.8$  Hz, 1H), 4.19 – 4.09 (m, 2H), 2.95 (s, 6H), 1.44 (s, 18H), 1.08 (t,  $J = 7.1$  Hz, 3H).

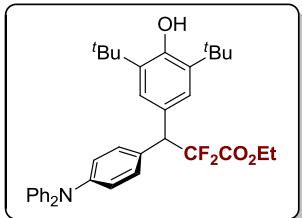
$^{13}\text{C}$  NMR (125 MHz,  $\text{CDCl}_3$ )  $\delta$  164.3 (t,  $J = 32.8$  Hz), 153.1, 149.9, 135.6, 130.3, 126.6, 126.2, 123.6, 116.4 (t,  $J = 255.4$  Hz),

112.4, 62.3, 54.7 (t,  $J = 21.8$  Hz), 40.5, 34.3, 30.3, 13.7.

$^{19}\text{F}$  NMR (471 MHz,  $\text{CDCl}_3$ )  $\delta$  -106.26 (d,  $J = 253.5$  Hz), -107.05 (d,  $J = 253.5$  Hz).

HRMS: (ESI $^+$ ) [M+Na] $^+$  calcd for  $\text{C}_{27}\text{H}_{37}\text{F}_2\text{NO}_3\text{Na}$ : 484.2634, found: 484.2640.

**Ethyl 3-(3,5-di-*tert*-butyl-4-hydroxyphenyl)-3-(4-(diphenylamino)phenyl)-2,2-difluoropropanoate (3o)**



$^1\text{H}$  NMR (500 MHz,  $\text{CDCl}_3$ )  $\delta$  7.34 – 7.29 (m, 2H), 7.28 – 7.24 (m, 4H), 7.22 (s, 2H), 7.12 – 7.07 (m, 4H), 7.06 – 7.01 (m, 4H), 5.21 (s, 1H), 4.60 (t,  $J = 18.5$  Hz, 1H), 4.23 – 4.10 (m, 2H), 1.44 (s, 18H), 1.09 (t,  $J = 7.1$  Hz, 3H).

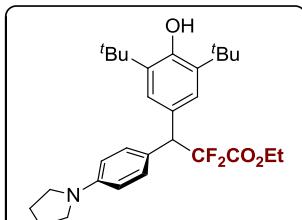
$^{13}\text{C}$  NMR (125 MHz,  $\text{CDCl}_3$ )  $\delta$  164.2 (t,  $J = 32.6$  Hz), 153.3, 147.6, 147.2, 135.8, 130.4, 129.71, 129.68, 129.2, 126.3, 126.09,

126.06, 124.5, 123.3, 122.9, 116.2 (t,  $J = 255.8$  Hz), 62.5, 54.9 (t,  $J = 21.8$  Hz), 34.4, 30.3, 13.7.

$^{19}\text{F}$  NMR (471 MHz,  $\text{CDCl}_3$ )  $\delta$  -105.95 (d,  $J = 254.1$  Hz), -107.43 (d,  $J = 254.1$  Hz).

HRMS: (ESI $^+$ ) [M+Na] $^+$  calcd for  $\text{C}_{37}\text{H}_{41}\text{F}_2\text{NO}_3\text{Na}$ : 608.2947, found: 608.2950.

**Ethyl 3-(3,5-di-*tert*-butyl-4-hydroxyphenyl)-2,2-difluoro-3-(4-(pyrrolidin-1-yl)phenyl)-propanoate (3p)**



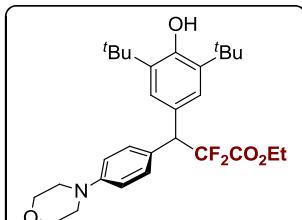
$^1\text{H}$  NMR (500 MHz,  $\text{CDCl}_3$ )  $\delta$  7.29 (d,  $J = 8.7$  Hz, 2H), 7.23 (s, 2H), 6.54 (d,  $J = 8.7$  Hz, 2H), 5.15 (s, 1H), 4.55 (t,  $J = 18.8$  Hz, 1H), 4.22 – 4.05 (m, 2H), 3.34 – 3.21 (m, 4H), 2.11 – 1.91 (m, 4H), 1.44 (s, 18H), 1.09 (t,  $J = 7.1$  Hz, 3H).

$^{13}\text{C}$  NMR (125 MHz,  $\text{CDCl}_3$ )  $\delta$  164.4 (t,  $J = 32.8$  Hz), 153.0, 147.3, 135.5, 130.3, 126.8, 126.2, 122.4, 116.5 (t,  $J = 255.1$  Hz), 111.5, 62.3, 54.7 (t,  $J = 21.7$  Hz), 47.5, 34.3, 30.3, 25.4, 13.7.

$^{19}\text{F}$  NMR (471 MHz,  $\text{CDCl}_3$ )  $\delta$  -106.26 (d,  $J = 254.3$  Hz), -106.93 (d,  $J = 254.3$  Hz).

HRMS: (ESI $^+$ ) [M+Na] $^+$  calcd for  $\text{C}_{29}\text{H}_{39}\text{F}_2\text{NO}_4\text{Na}$ : 510.2790, found: 510.2794.

**Ethyl 3-(3,5-di-*tert*-butyl-4-hydroxyphenyl)-2,2-difluoro-3-(4-morpholinophenyl)-propanoate (3q)**



$^1\text{H}$  NMR (500 MHz,  $\text{CDCl}_3$ )  $\delta$  7.35 (d,  $J = 8.8$  Hz, 2H), 7.20 (s, 2H), 6.89 (d,  $J = 8.8$  Hz, 2H), 5.17 (s, 1H), 4.58 (t,  $J = 18.6$  Hz, 1H), 4.20 – 4.08 (m, 2H), 3.97 – 3.80 (m, 4H), 3.23 – 3.10 (m, 4H), 1.43 (s, 18H), 1.07 (t,  $J = 7.1$  Hz, 3H).

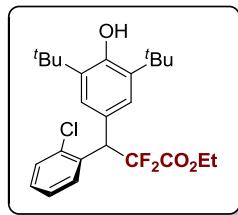
$^{13}\text{C}$  NMR (125 MHz,  $\text{CDCl}_3$ )  $\delta$  164.2 (t,  $J = 32.8$  Hz), 153.2, 150.5, 135.7, 130.4, 127.29, 127.28, 126.32, 126.30, 126.2, 116.3 (t,

$J = 257.1$  Hz), 115.4, 66.9, 62.4, 54.7 (t,  $J = 21.5$  Hz), 49.1, 34.3, 30.3, 13.7.

$^{19}\text{F}$  NMR (471 MHz,  $\text{CDCl}_3$ )  $\delta$  -106.00 (d,  $J = 254.2$  Hz), -107.26 (d,  $J = 254.2$  Hz).

HRMS: (ESI $^+$ ) [M+Na] $^+$  calcd for  $\text{C}_{29}\text{H}_{39}\text{F}_2\text{NO}_4\text{Na}$ : 526.2739, found: 526.2742.

**Ethyl 3-(2-chlorophenyl)-3-(3,5-di-*tert*-butyl-4-hydroxyphenyl)-2,2-difluoropropionate (3r)**

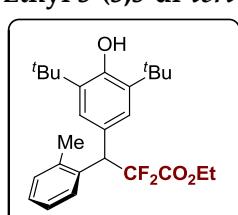


$^1\text{H}$  NMR (500 MHz,  $\text{CDCl}_3$ )  $\delta$  7.75 (dd,  $J = 7.9, 1.5$  Hz, 1H), 7.40 (dd,  $J = 7.9, 1.5$  Hz, 1H), 7.31 (td,  $J = 7.7, 1.5$  Hz, 1H), 7.24 (dd,  $J = 7.7, 1.7$  Hz, 1H), 7.21 (s, 2H), 5.36 (dd,  $J = 19.4, 17.6$  Hz, 1H), 5.21 (s, 1H), 4.17 (q,  $J = 7.1$  Hz, 2H), 1.42 (s, 18H), 1.09 (t,  $J = 7.1$  Hz, 3H).  
 $^{13}\text{C}$  NMR (125 MHz,  $\text{CDCl}_3$ )  $\delta$  163.7 (t,  $J = 32.5$  Hz), 153.4, 135.7, 134.8, 134.37, 134.33, 129.9, 129.7, 128.7, 126.9, 126.6, 124.62, 124.58, 116.0 (t,  $J = 256.2$  Hz), 62.7, 50.5 (t,  $J = 22.1$  Hz), 34.3, 30.2, 13.6.

$^{19}\text{F}$  NMR (471 MHz,  $\text{CDCl}_3$ )  $\delta$  -103.78 (d,  $J = 257.0$  Hz), -107.27 (d,  $J = 257.0$  Hz).

HRMS: (ESI $^+$ ) [M+Na] $^+$  calcd for  $\text{C}_{28}\text{H}_{31}\text{ClF}_2\text{O}_3\text{Na}$ : 475.1822, found: 475.1823.

**Ethyl 3-(3,5-di-*tert*-butyl-4-hydroxyphenyl)-2,2-difluoro-3-(o-tolyl)propanoate (3s)**

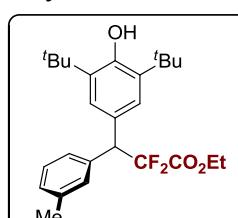


$^1\text{H}$  NMR (500 MHz,  $\text{CDCl}_3$ )  $\delta$  7.64 (d,  $J = 7.7$  Hz, 1H), 7.24 (td,  $J = 7.4, 2.1$  Hz, 1H), 7.21 – 7.14 (m, 2H), 7.17 (s, 2H), 5.17 (s, 1H), 4.91 (dd,  $J = 20.7, 16.8$  Hz, 1H), 4.13 (q,  $J = 7.1$  Hz, 2H), 2.37 (s, 3H), 1.41 (s, 18H), 1.05 (t,  $J = 7.1$  Hz, 3H).  
 $^{13}\text{C}$  NMR (125 MHz,  $\text{CDCl}_3$ )  $\delta$  164.2 (t,  $J = 32.6$  Hz), 153.3, 136.8, 135.6, 134.90, 134.85, 130.7, 128.0, 127.4, 126.8, 126.1, 125.10, 125.07, 116.3 (t,  $J = 257.6$  Hz), 62.5, 50.5 (t,  $J = 22.1$  Hz), 34.3, 30.3, 20.2, 13.5.

$^{19}\text{F}$  NMR (471 MHz,  $\text{CDCl}_3$ )  $\delta$  -101.76 (d,  $J = 256.6$  Hz), -108.33 (d,  $J = 256.6$  Hz).

HRMS: (ESI $^+$ ) [M+Na] $^+$  calcd for  $\text{C}_{26}\text{H}_{34}\text{F}_2\text{O}_3\text{Na}$ : 455.2368, found: 455.2369.

**Ethyl 3-(3,5-di-*tert*-butyl-4-hydroxyphenyl)-2,2-difluoro-3-(m-tolyl)propanoate (3t)**

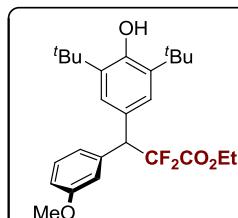


$^1\text{H}$  NMR (500 MHz,  $\text{CDCl}_3$ )  $\delta$  7.64 (d,  $J = 7.7$  Hz, 1H), 7.24 (td,  $J = 7.4, 2.1$  Hz, 1H), 7.20 – 7.14 (m, 2H), 7.17 (s, 2H), 5.17 (s, 1H), 4.92 (dd,  $J = 20.6, 16.8$  Hz, 1H), 4.13 (q,  $J = 7.1$  Hz, 2H), 2.37 (s, 3H), 1.42 (s, 18H), 1.05 (t,  $J = 7.1$  Hz, 3H).  
 $^{13}\text{C}$  NMR (125 MHz,  $\text{CDCl}_3$ )  $\delta$  164.2 (t,  $J = 32.6$  Hz), 153.3, 136.8, 135.6, 134.90, 134.85, 130.7, 128.03, 128.00, 127.4, 126.7, 126.1, 125.1, 116.33 (d,  $J = 255.1$  Hz), 62.5, 50.5 (t,  $J = 21.9$  Hz), 34.3, 30.3, 20.2, 13.5.

$^{19}\text{F}$  NMR (471 MHz,  $\text{CDCl}_3$ )  $\delta$  -101.80 (d,  $J = 255.9$  Hz), -108.30 (d,  $J = 255.9$  Hz).

HRMS: (ESI $^+$ ) [M+H] $^+$  calcd for  $\text{C}_{26}\text{H}_{35}\text{F}_2\text{O}_3$ : 433.2549, found: 433.2540.

**Ethyl 3-(3,5-di-*tert*-butyl-4-hydroxyphenyl)-2,2-difluoro-3-(3-methoxyphenyl)propanoate (3u)**

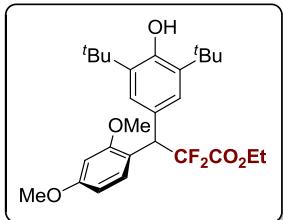


$^1\text{H}$  NMR (500 MHz,  $\text{CDCl}_3$ )  $\delta$  7.31 – 7.24 (m, 1H), 7.22 (s, 2H), 7.06 (d,  $J = 7.6$  Hz, 1H), 7.00 (t,  $J = 2.1$  Hz, 1H), 6.85 (dd,  $J = 8.2, 2.5$  Hz, 1H), 5.20 (s, 1H), 4.64 (t,  $J = 18.3$  Hz, 1H), 4.15 (qd,  $J = 7.1, 5.3$  Hz, 2H), 3.82 (s, 3H), 1.44 (s, 18H), 1.07 (t,  $J = 7.1$  Hz, 3H).  
 $^{13}\text{C}$  NMR (125 MHz,  $\text{CDCl}_3$ )  $\delta$  164.0 (t,  $J = 32.6$  Hz), 159.5, 153.4, 137.58, 137.55, 135.8, 129.4, 126.3, 125.78, 125.75, 121.9, 116.1 (t,  $J = 256.0$  Hz), 115.5, 113.0, 62.5, 55.4 (t,  $J = 21.8$  Hz), 55.1, 34.3, 30.2, 13.6.

$^{19}\text{F}$  NMR (471 MHz,  $\text{CDCl}_3$ )  $\delta$  -105.59 (d,  $J = 255.3$  Hz), -107.10 (d,  $J = 255.3$  Hz).

**HRMS:** (ESI<sup>+</sup>) [M+Na]<sup>+</sup> calcd for C<sub>26</sub>H<sub>34</sub>F<sub>2</sub>O<sub>4</sub>Na: 471.2317, found: 471.2313.

**Ethyl 3-(3,5-di-*tert*-butyl-4-hydroxyphenyl)-3-(2,4-dimethoxyphenyl)-2,2-difluoropropanoate (3v)**



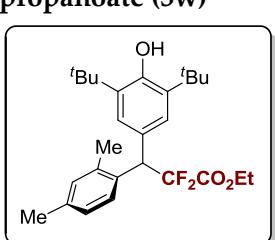
**<sup>1</sup>H NMR** (500 MHz, CDCl<sub>3</sub>) δ 7.54 (d, *J* = 8.6 Hz, 1H), 7.23 (s, 2H), 6.53 (dd, *J* = 8.6, 2.5 Hz, 1H), 6.45 (d, *J* = 2.4 Hz, 1H), 5.24 (dd, *J* = 21.1, 17.7 Hz, 1H), 5.15 (s, 1H), 4.14 (q, *J* = 7.1 Hz, 2H), 3.81 (s, 3H), 3.81 (s, 3H), 1.43 (s, 18H), 1.08 (t, *J* = 7.1 Hz, 3H).

**<sup>13</sup>C NMR** (125 MHz, CDCl<sub>3</sub>) δ 164.3 (t, *J* = 32.8 Hz), 160.1, 158.1, 153.1, 135.4, 130.0, 126.5, 126.25, 126.22, 117.39, 117.35, 116.6 (t, *J* = 254.7 Hz), 104.3, 98.7, 62.3, 55.7, 55.2, 45.7 (t, *J* = 22.0 Hz), 34.3, 30.3, 13.6.

**<sup>19</sup>F NMR** (471 MHz, CDCl<sub>3</sub>) δ -103.43 (d, *J* = 252.9 Hz), -107.79 (d, *J* = 252.9 Hz).

**HRMS:** (ESI<sup>+</sup>) [M+Na]<sup>+</sup> calcd for C<sub>27</sub>H<sub>36</sub>F<sub>2</sub>O<sub>5</sub>Na: 501.2423, found: 501.2419.

**Ethyl 3-(3,5-di-*tert*-butyl-4-hydroxyphenyl)-3-(2,4-dimethylphenyl)-2,2-difluoropropanoate (3w)**



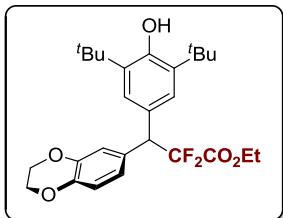
**<sup>1</sup>H NMR** (500 MHz, CDCl<sub>3</sub>) δ 7.52 (d, *J* = 7.7 Hz, 1H), 7.17 (s, 2H), 7.10 – 7.01 (m, 1H), 6.98 (d, *J* = 1.8 Hz, 1H), 5.16 (s, 1H), 4.87 (dd, *J* = 20.6, 17.1 Hz, 1H), 4.13 (qd, *J* = 7.1, 1.6 Hz, 2H), 2.33 (s, 3H), 2.30 (s, 3H), 1.42 (s, 18H), 1.07 (t, *J* = 7.1 Hz, 3H).

**<sup>13</sup>C NMR** (125 MHz, CDCl<sub>3</sub>) δ 164.2 (t, *J* = 32.7 Hz), 153.2, 136.9, 136.5, 135.5, 131.81, 131.77, 131.5, 128.0, 126.8, 126.7, 125.42, 125.39, 116.4 (dd, *J* = 257.0, 252.8 Hz), 62.5, 50.2 (t, *J* = 21.8 Hz), 34.3, 30.2, 20.9, 20.1, 13.8.

**<sup>19</sup>F NMR** (471 MHz, CDCl<sub>3</sub>) δ -102.10 (d, *J* = 255.8 Hz), -108.20 (d, *J* = 255.8 Hz).

**HRMS:** (ESI<sup>+</sup>) [M+Na]<sup>+</sup> calcd for C<sub>27</sub>H<sub>36</sub>F<sub>2</sub>O<sub>3</sub>Na: 469.2525, found: 469.2529.

**Ethyl 3-(3,5-di-*tert*-butyl-4-hydroxyphenyl)-3-(2,3-dihydrobenzo[b][1,4]dioxin-7-yl)-2,2-difluoropropanoate (3x)**



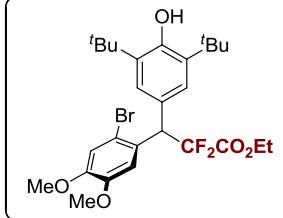
**<sup>1</sup>H NMR** (500 MHz, CDCl<sub>3</sub>) δ 7.19 (s, 2H), 6.98 – 6.89 (m, 2H), 6.84 (d, *J* = 8.3 Hz, 1H), 5.19 (s, 1H), 4.55 (t, *J* = 18.4 Hz, 1H), 4.25 (s, 4H), 4.14 (qt, *J* = 10.1, 6.9 Hz, 2H), 1.44 (s, 18H), 1.08 (t, *J* = 7.1 Hz, 3H).

**<sup>13</sup>C NMR** (125 MHz, CDCl<sub>3</sub>) δ 164.1 (t, *J* = 32.7 Hz), 153.3, 143.3, 143.0, 135.8, 129.28, 129.25, 128.08, 128.07, 126.1, 122.6, 118.6, 117.1, 116.1 (t, *J* = 255.6 Hz), 64.31, 64.27, 62.5, 54.8 (t, *J* = 21.8 Hz), 34.3, 30.2, 13.6.

**<sup>19</sup>F NMR** (471 MHz, CDCl<sub>3</sub>) δ -105.37 (d, *J* = 254.9 Hz), -107.83 (d, *J* = 254.8 Hz).

**HRMS:** (ESI<sup>+</sup>) [M+Na]<sup>+</sup> calcd for C<sub>27</sub>H<sub>34</sub>F<sub>2</sub>O<sub>5</sub>Na: 499.2267, found: 499.2262.

**Ethyl 3-(2-bromo-4,5-dimethoxyphenyl)-3-(3,5-di-*tert*-butyl-4-hydroxyphenyl)-2,2-difluoropropanoate (3y)**



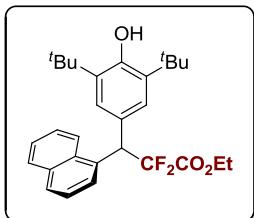
**<sup>1</sup>H NMR** (500 MHz, CDCl<sub>3</sub>) δ 7.27 (s, 1H), 7.24 (s, 2H), 7.04 (s, 1H), 5.30 – 5.18 (m, 2H), 5.20 (s, 1H), 4.18 (q, *J* = 7.1 Hz, 2H), 3.91 (s, 3H), 3.86 (s, 3H), 1.43 (s, 18H), 1.13 (t, *J* = 7.1 Hz, 3H).

**<sup>13</sup>C NMR** (125 MHz, CDCl<sub>3</sub>) δ 163.7 (t, *J* = 32.6 Hz), 153.4, 148.8, 148.3, 135.7, 127.79, 127.74, 126.4, 124.96, 124.93, 116.1 (t, *J* = 255.3 Hz), 115.8, 115.6, 112.3, 62.3, 56.0, 55.9, 53.0 (t, *J* = 22.0 Hz), 34.3, 30.2, 13.7.

**<sup>19</sup>F NMR** (471 MHz, CDCl<sub>3</sub>) δ -103.43 (d, *J* = 256.2 Hz), -107.85 (d, *J* = 256.2 Hz).

**HRMS:** (ESI<sup>+</sup>) [M+Na]<sup>+</sup> calcd for C<sub>27</sub>H<sub>35</sub>BrF<sub>2</sub>O<sub>5</sub>Na: 579.1528, found: 579.1533.

**Ethyl 3-(3,5-di-tert-butyl-4-hydroxyphenyl)-2,2-difluoro-3-(naphthalen-1-yl)propanoate (3z)**



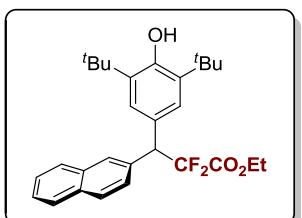
**<sup>1</sup>H NMR** (500 MHz, CDCl<sub>3</sub>) δ 8.15 (d, *J* = 8.5 Hz, 1H), 7.90 – 7.78 (m, 3H), 7.57 – 7.46 (m, 3H), 7.27 (s, 2H), 5.58 (dd, *J* = 20.0, 16.6 Hz, 1H), 5.17 (s, 1H), 4.06 (q, *J* = 7.1 Hz, 2H), 1.40 (s, 18H), 0.90 (t, *J* = 7.1 Hz, 3H).

**<sup>13</sup>C NMR** (125 MHz, CDCl<sub>3</sub>) δ 164.1 (t, *J* = 32.6 Hz), 153.4, 135.6, 134.1, 132.26, 132.21, 131.9, 128.9, 128.3, 126.6, 126.4, 126.23, 126.19, 125.5, 125.4, 125.2, 123.3, 116.3 (dd, *J* = 257.0, 252.0 Hz), 62.6, 49.7 (t, *J* = 21.8 Hz), 34.3, 30.2, 13.5.

**<sup>19</sup>F NMR** (471 MHz, CDCl<sub>3</sub>) δ -101.26 (d, *J* = 256.9 Hz), -107.67 (d, *J* = 256.9 Hz).

**HRMS:** (ESI<sup>+</sup>) [M+Na]<sup>+</sup> calcd for C<sub>29</sub>H<sub>34</sub>F<sub>2</sub>O<sub>3</sub>Na: 491.2368, found: 491.2374.

**Ethyl 3-(3,5-di-tert-butyl-4-hydroxyphenyl)-2,2-difluoro-3-(naphthalen-2-yl)propanoate (3aa)**



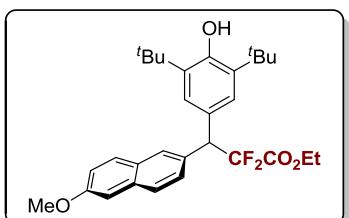
**<sup>1</sup>H NMR** (500 MHz, CDCl<sub>3</sub>) δ 7.92 (d, *J* = 1.8 Hz, 1H), 7.88 – 7.79 (m, 3H), 7.57 (dd, *J* = 8.6, 1.7 Hz, 1H), 7.54 – 7.46 (m, 2H), 7.27 (s, 2H), 5.20 (s, 1H), 4.85 (t, *J* = 18.3 Hz, 1H), 4.18 – 4.09 (m, 2H), 1.43 (s, 18H), 1.04 (t, *J* = 7.1 Hz, 3H).

**<sup>13</sup>C NMR** (125 MHz, CDCl<sub>3</sub>) δ 164.1 (t, *J* = 32.5 Hz), 153.4, 135.8, 133.71, 133.68, 133.3, 132.6, 128.6, 128.1, 128.0, 127.5, 127.4, 126.4, 126.11, 126.07, 125.90, 125.88, 116.2 (t, *J* = 256.0 Hz), 62.6, 55.5 (t, *J* = 21.8 Hz), 34.4, 30.3, 13.6.

**<sup>19</sup>F NMR** (471 MHz, CDCl<sub>3</sub>) δ -105.72 (d, *J* = 255.9 Hz), -106.50 (d, *J* = 255.9 Hz).

**HRMS:** (ESI<sup>+</sup>) [M+Na]<sup>+</sup> calcd for C<sub>29</sub>H<sub>34</sub>F<sub>2</sub>O<sub>3</sub>Na: 491.2368, found: 491.2372.

**Ethyl 3-(3,5-di-tert-butyl-4-hydroxyphenyl)-2,2-difluoro-3-(6-methoxynaphthalen-2-yl)propanoate (3ab)**



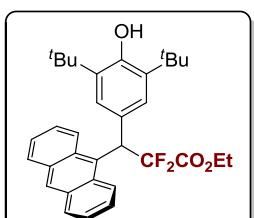
**<sup>1</sup>H NMR** (500 MHz, CDCl<sub>3</sub>) δ 7.87 – 7.80 (m, 1H), 7.73 (dd, *J* = 11.1, 8.8 Hz, 2H), 7.52 (dd, *J* = 8.5, 1.8 Hz, 1H), 7.26 (s, 2H), 7.16 (dd, *J* = 8.9, 2.5 Hz, 1H), 7.12 (d, *J* = 2.5 Hz, 1H), 5.19 (s, 1H), 4.81 (t, *J* = 18.4 Hz, 1H), 4.12 (qd, *J* = 7.1, 5.2 Hz, 2H), 3.94 (s, 3H), 1.43 (s, 18H), 1.04 (t, *J* = 7.1 Hz, 3H).

**<sup>13</sup>C NMR** (125 MHz, CDCl<sub>3</sub>) δ 164.1 (t, *J* = 32.8 Hz), 157.9, 153.3, 135.8, 133.8, 131.3, 129.5, 128.7, 128.4, 127.9, 127.0, 126.4, 126.1, 119.0, 116.3 (t, *J* = 256.4 Hz), 105.5, 62.5, 55.4 (t, *J* = 21.9 Hz), 55.3, 34.3, 30.3, 13.6.

**<sup>19</sup>F NMR** (471 MHz, CDCl<sub>3</sub>) δ -103.51 (d, *J* = 257.1 Hz), -109.00 (d, *J* = 257.1 Hz).

**HRMS:** (ESI<sup>+</sup>) [M+Na]<sup>+</sup> calcd for C<sub>30</sub>H<sub>36</sub>F<sub>2</sub>O<sub>4</sub>Na: 521.2474, found: 521.2479.

**Ethyl 3-(anthracen-9-yl)-3-(3,5-di-tert-butyl-4-hydroxyphenyl)-2,2-difluoropropanoate (3ac)**



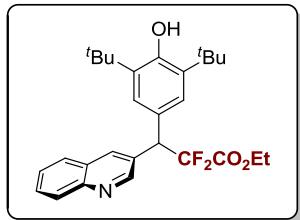
**<sup>1</sup>H NMR** (500 MHz, CDCl<sub>3</sub>) δ 7.82 (dd, *J* = 8.1, 1.3 Hz, 1H), 7.46 – 7.37 (m, 4H), 7.31 – 7.27 (m, 1H), 7.24 (s, 2H), 7.21 (td, *J* = 7.6, 1.3 Hz, 1H), 7.09 (td, *J* = 7.6, 1.3 Hz, 1H), 7.06 (s, 1H), 5.24 (s, 1H), 4.74 (dd, *J* = 17.3, 14.0 Hz, 1H), 4.29 – 4.14 (m, 2H), 1.37 (s, 18H), 1.23 (t, *J* = 7.2 Hz, 3H).

**<sup>13</sup>C NMR** (125 MHz, CDCl<sub>3</sub>) δ 164.1 (t, *J* = 32.1 Hz), 153.2, 140.9, 137.0, 135.5, 133.5, 131.24, 131.21, 129.63, 129.56, 129.2, 128.94, 128.92, 128.3, 128.2, 128.0, 126.8, 126.7, 126.6, 126.3, 123.6, 116.3 (t, *J* = 258.9 Hz), 62.8, 51.2 (t, *J* = 23.6 Hz), 34.2, 30.2, 13.8.

**<sup>19</sup>F NMR** (471 MHz, CDCl<sub>3</sub>) δ -106.98 (d, *J* = 245.3 Hz), -109.52 (d, *J* = 245.3 Hz).

**HRMS:** (ESI<sup>+</sup>) [M+Na]<sup>+</sup> calcd for C<sub>33</sub>H<sub>36</sub>F<sub>2</sub>O<sub>3</sub>Na: 541.2525, found: 541.2529.

**Ethyl 3-(3,5-di-*tert*-butyl-4-hydroxyphenyl)-2,2-difluoro-3-(quinoline-2-yl)propanoate (3ad)**



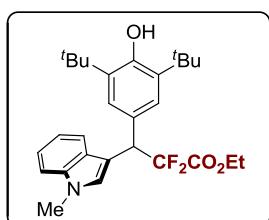
**<sup>1</sup>H NMR** (500 MHz, CDCl<sub>3</sub>) δ 8.94 (d, *J* = 2.1 Hz, 1H), 8.26 (d, *J* = 2.1 Hz, 1H), 8.10 (d, *J* = 8.4 Hz, 1H), 7.86 (dd, *J* = 8.2, 1.5 Hz, 1H), 7.73 (ddd, *J* = 8.3, 6.8, 1.4 Hz, 1H), 7.58 (ddd, *J* = 8.0, 6.8, 1.1 Hz, 1H), 7.24 (s, 2H), 5.26 (s, 1H), 4.90 (dd, *J* = 20.3, 15.7 Hz, 1H), 4.21 – 4.12 (m, 2H), 1.43 (s, 18H), 1.07 (t, *J* = 7.1 Hz, 3H).

**<sup>13</sup>C NMR** (125 MHz, CDCl<sub>3</sub>) δ 163.6 (t, *J* = 32.3 Hz), 153.7, 151.8, 147.3, 136.3, 136.1, 129.65, 129.32, 129.29, 129.1, 128.0, 127.6, 126.9, 126.2, 124.80, 124.75, 115.9 (t, *J* = 256.1 Hz), 62.8, 53.3 (t, *J* = 22.2 Hz), 34.4, 30.2, 13.6.

**<sup>19</sup>F NMR** (471 MHz, CDCl<sub>3</sub>) δ -103.39 (d, *J* = 257.6 Hz), -108.54 (d, *J* = 257.6 Hz).

**HRMS:** (ESI<sup>+</sup>) [M+Na]<sup>+</sup> calcd for C<sub>28</sub>H<sub>33</sub>F<sub>2</sub>NO<sub>3</sub>Na: 492.2321, found: 492.2319.

**Ethyl 3-(3,5-di-*tert*-butyl-4-hydroxyphenyl)-2,2-difluoro-3-(1-methyl-1H-indol-3-yl)propanoate (3ae)**



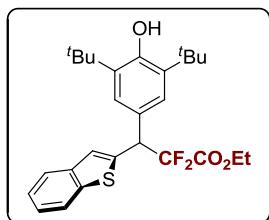
**<sup>1</sup>H NMR** (500 MHz, CDCl<sub>3</sub>) δ 7.62 (d, *J* = 8.0 Hz, 1H), 7.31 (dt, *J* = 8.2, 1.0 Hz, 1H), 7.28 (s, 2H), 7.26 – 7.21 (m, 2H), 7.13 (ddd, *J* = 7.9, 7.0, 1.1 Hz, 1H), 5.15 (s, 1H), 4.99 (dd, *J* = 20.4, 16.1 Hz, 1H), 4.08 (qd, *J* = 7.1, 5.3 Hz, 2H), 3.81 (s, 3H), 1.43 (s, 18H), 1.01 (t, *J* = 7.1 Hz, 3H).

**<sup>13</sup>C NMR** (125 MHz, CDCl<sub>3</sub>) δ 164.3 (t, *J* = 32.8 Hz), 153.2, 136.6, 135.6, 128.0, 127.7, 126.3, 126.23, 126.20, 121.8, 119.2, 119.1, 116.4 (t, *J* = 255.1 Hz), 109.32, 109.27, 109.2, 62.4, 46.9 (t, *J* = 22.8 Hz), 34.3, 32.9, 30.3, 13.6.

**<sup>19</sup>F NMR** (471 MHz, CDCl<sub>3</sub>) δ -106.05 (d, *J* = 250.9 Hz), -109.13 (d, *J* = 250.9 Hz).

**HRMS:** (ESI<sup>+</sup>) [M+Na]<sup>+</sup> calcd for C<sub>28</sub>H<sub>35</sub>F<sub>2</sub>NO<sub>3</sub>Na: 494.2477, found: 494.2486.

**Ethyl 3-(benzo[b]thiophen-2-yl)-3-(3,5-di-*tert*-butyl-4-hydroxyphenyl)-2,2-difluoro-propanoate (3af)**



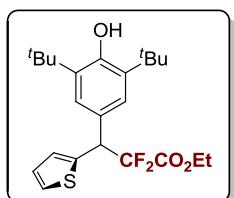
**<sup>1</sup>H NMR** (500 MHz, CDCl<sub>3</sub>) δ 7.88 (dd, *J* = 7.1, 1.5 Hz, 1H), 7.77 (dd, *J* = 7.4, 1.6 Hz, 1H), 7.69 (d, *J* = 1.4 Hz, 1H), 7.42 – 7.33 (m, 2H), 7.24 (s, 2H), 5.20 (s, 1H), 5.14 (dd, *J* = 19.7, 15.4 Hz, 1H), 4.13 (qd, *J* = 7.1, 2.1 Hz, 2H), 1.42 (s, 18H), 1.02 (t, *J* = 7.1 Hz, 3H).

**<sup>13</sup>C NMR** (125 MHz, CDCl<sub>3</sub>) δ 164.0 (t, *J* = 32.5 Hz), 153.6, 139.8, 138.6, 135.7, 130.18, 130.14, 126.4, 124.4, 124.18, 124.15, 124.1, 122.7, 121.7, 117.9, 115.9 (t, *J* = 255.5 Hz), 62.7, 48.8 (t, *J* = 22.7 Hz), 34.3, 30.2, 13.6.

**<sup>19</sup>F NMR** (471 MHz, CDCl<sub>3</sub>) δ -104.66 (d, *J* = 255.0 Hz), -109.15 (d, *J* = 255.0 Hz).

**HRMS:** (ESI<sup>+</sup>) [M+Na]<sup>+</sup> calcd for C<sub>27</sub>H<sub>32</sub>F<sub>2</sub>O<sub>3</sub>SnA: 497.1932, found: 497.1935.

**Ethyl 3-(3,5-di-*tert*-butyl-4-hydroxyphenyl)-2,2-difluoro-3-(thiophen-2-yl)propanoate (3ag)**



**<sup>1</sup>H NMR** (500 MHz, CDCl<sub>3</sub>) δ 7.31 – 7.27 (m, 1H), 7.26 (s, 2H), 7.12 (d, *J* = 3.5 Hz, 1H), 7.00 (dd, *J* = 5.1, 3.5 Hz, 1H), 5.24 (s, 1H), 4.92 (t, *J*

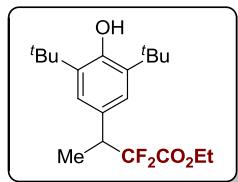
= 17.4 Hz, 1H), 4.15 (qd,  $J$  = 7.1, 2.6 Hz, 2H), 1.44 (s, 18H), 1.10 (t,  $J$  = 7.1 Hz, 3H).

**$^{13}\text{C}$  NMR** (125 MHz,  $\text{CDCl}_3$ )  $\delta$  163.8 (t,  $J$  = 32.3 Hz), 153.7, 138.01, 137.97, 135.9, 127.6, 126.7, 126.3, 125.6, 125.28, 125.24, 115.4 (t,  $J$  = 257.2 Hz), 62.6, 51.1 (t,  $J$  = 22.8 Hz), 34.4, 30.2, 13.7.

**$^{19}\text{F}$  NMR** (471 MHz,  $\text{CDCl}_3$ )  $\delta$  -106.54 (d,  $J$  = 251.3 Hz), -109.76 (d,  $J$  = 251.3 Hz).

**HRMS:** (ESI $^+$ ) [M+Na] $^+$  calcd for  $\text{C}_{23}\text{H}_{30}\text{F}_2\text{O}_3\text{SNa}$ : 447.1776, found: 447.1779.

**Ethyl 3-(3,5-di-*tert*-butyl-4-hydroxyphenyl)-2,2-difluorobutanoate (3ah)**



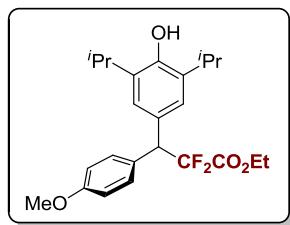
**$^1\text{H}$  NMR** (500 MHz,  $\text{CDCl}_3$ )  $\delta$  7.07 (s, 2H), 5.20 (s, 1H), 4.21 – 4.11 (m, 2H), 3.43 (ddq,  $J$  = 20.5, 14.3, 7.2 Hz, 1H), 1.47 (d,  $J$  = 9.6 Hz, 3H), 1.45 (s, 18H), 1.15 (t,  $J$  = 7.1 Hz, 3H).

**$^{13}\text{C}$  NMR** (125 MHz,  $\text{CDCl}_3$ )  $\delta$  164.3 (t,  $J$  = 32.9 Hz), 153.3, 135.7, 127.3, 125.4, 116.9 (dd,  $J$  = 257.0, 253.4 Hz), 62.3, 44.2 (t,  $J$  = 22.5 Hz), 34.3, 30.3, 13.79 (t,  $J$  = 2.5 Hz), 13.76.

**$^{19}\text{F}$  NMR** (471 MHz,  $\text{CDCl}_3$ )  $\delta$  -108.55 (d,  $J$  = 248.6 Hz), -115.40 (d,  $J$  = 248.6 Hz).

**HRMS:** (ESI $^+$ ) [M+Na] $^+$  calcd for  $\text{C}_{20}\text{H}_{30}\text{F}_2\text{O}_3\text{Na}$ : 379.2055, found: 379.2054.

**Ethyl 2,2-difluoro-3-(4-hydroxy-3,5-diisopropylphenyl)-3-(4-methoxyphenyl)-propanoate (3ai)**



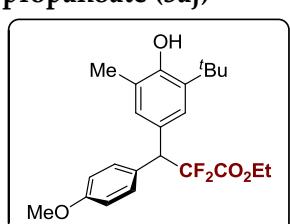
**$^1\text{H}$  NMR** (500 MHz,  $\text{CDCl}_3$ )  $\delta$  7.34 (d,  $J$  = 8.7 Hz, 2H), 7.07 (s, 2H), 6.88 (d,  $J$  = 8.7 Hz, 2H), 4.78 (s, 1H), 4.64 (t,  $J$  = 18.4 Hz, 1H), 4.14 (qd,  $J$  = 7.1, 4.4 Hz, 2H), 3.81 (s, 3H), 3.13 (p,  $J$  = 6.9 Hz, 2H), 1.26 (d,  $J$  = 6.9 Hz, 12H), 1.10 (t,  $J$  = 7.1 Hz, 3H).

**$^{13}\text{C}$  NMR** (125 MHz,  $\text{CDCl}_3$ )  $\delta$  164.1 (t,  $J$  = 32.8 Hz), 159.0, 149.5, 133.60, 130.7, 128.1, 128.0, 127.52, 127.49, 124.7, 116.1 (t,  $J$  = 255.8 Hz), 113.8, 62.5, 55.2, 54.5 (t,  $J$  = 21.8 Hz), 27.2, 22.7, 22.6, 13.7.

**$^{19}\text{F}$  NMR** (471 MHz,  $\text{CDCl}_3$ )  $\delta$  -105.62 (d,  $J$  = 255.5 Hz), -107.44 (d,  $J$  = 255.5 Hz).

**HRMS:** (ESI $^+$ ) [M+Na] $^+$  calcd for  $\text{C}_{24}\text{H}_{30}\text{F}_2\text{O}_4\text{Na}$ : 443.2004, found: 443.2009.

**Ethyl 3-(3-(*tert*-butyl)-4-hydroxy-5-methylphenyl)-2,2-difluoro-3-(4-methoxyphenyl)-propanoate (3aj)**



**$^1\text{H}$  NMR** (500 MHz,  $\text{CDCl}_3$ )  $\delta$  7.35 (d,  $J$  = 8.7 Hz, 2H), 7.16 (d,  $J$  = 2.2 Hz, 1H), 7.04 (d,  $J$  = 2.1 Hz, 1H), 6.88 (d,  $J$  = 8.8 Hz, 2H), 4.77 (s, 1H), 4.61 (t,  $J$  = 18.5 Hz, 1H), 4.16 (qd,  $J$  = 7.1, 1.8 Hz, 2H), 3.81 (s, 3H), 2.23 (s, 3H), 1.40 (s, 9H), 1.12 (t,  $J$  = 7.1 Hz, 3H).

**$^{13}\text{C}$  NMR** (125 MHz,  $\text{CDCl}_3$ )  $\delta$  164.1 (t,  $J$  = 32.8 Hz), 159.0, 152.2, 135.5, 130.6, 129.4, 128.2, 126.9, 126.5, 123.1, 116.1 (t,  $J$  = 255.4 Hz), 113.9, 62.6, 55.2, 54.3 (t,  $J$  = 21.9 Hz), 34.5, 29.7, 16.1, 13.7.

**$^{19}\text{F}$  NMR** (471 MHz,  $\text{CDCl}_3$ )  $\delta$  -106.08 (d,  $J$  = 255.8 Hz), -107.11 (d,  $J$  = 255.8 Hz).

**HRMS:** (ESI $^+$ ) [M+Na] $^+$  calcd for  $\text{C}_{23}\text{H}_{28}\text{F}_2\text{O}_4\text{Na}$ : 429.1848, found: 429.1847.

#### IV. NMR Spectra

Please refer to next pages

