Supporting Information for Experiments

Cu/Chiral Phosphoric Acid-Catalyzed Asymmetric Three-Component Radical-Initiated 1,2-Dicarbofunctionalization of Alkenes

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Figure S2. X-ray of chiral compound 7G

Scheme S1. Asymmetric Intermolecular Three-Component Radical-Initiated Dicarbofunctionalization of *meta*-Phenol Substrate 1ba



General information

Most of reactions were carried out under argon atmosphere using Schlenk techniques. Reagents were purchased at the highest commercial quality and used without further purification, unless otherwise stated. CuI was purchased from Sigma-Aldrich. Chiral phosphoric acid (CPA) was purchased from Daicel Chiral Technologies (China). Analytical thin layer chromatography (TLC) was performed on precoated silica gel 60 GF254 plates. Flash column chromatography was performed using Tsingdao silica gel (60, particle size 0.040-0.063 mm). Visualization on TLC was achieved by use of UV light (254 nm) or iodine. NMR spectra were recorded on Bruker DRX-500 and DPX 400 spectrometer at 400 or 500 MHz for ¹H NMR, 101 or 126 MHz for ¹³C NMR and 376 MHz for ¹⁹F NMR in CDCl₃, or CD₃OD with tetramethylsilane (TMS) as internal standard. The chemical shifts are expressed in ppm and coupling constants are given in Hz. Data for ¹H NMR are recorded as follows: chemical shift (ppm), multiplicity (s, singlet; d, doublet; t, triplet; q, quarter; p, pentet, m, multiplet; br, broad), coupling constant (Hz), integration. Data for ¹³C NMR are reported in terms of chemical shift (δ, ppm) . Mass spectrometric data were obtained using Bruker Apex IV RTMS. Enantiomeric excess (ee) was determined using Agilent high-performance liquid chromatography (HPLC) with a Hatachi detector ($\lambda = 254$ or 214 nm). Column conditions are reported in the experimental section below. Absolute configuration of a product was determined by X-ray analysis.

General procedure for the synthesis of substrates:

General synthesis of substrates 1a-1m and 1q



Under argon atmosphere, TIPSCl (250 mmol) was added dropwise to a solution of p-hydroxybenzaldehyde (200 mmol) and imidazole (500 mmol) in THF (250 mL). Then the reaction was stirred at 30 °C for 10 h. After complete conversion (monitored by TLC), the reaction mixture was extracted with EtOAc. The combined organic layers were concentrated *in vacuo*. The residue was purified by column chromatography on silica gel (petroleum ether /EtOAc = 80/1) to get **S-1a** (186 mmol, 93%).

Under argon atmosphere, CBr₄ (216 mmol) and PPh₃ (360 mmol) were added to a solution of **S-1a** (180 mmol) in DCM (250 mL) sequentially. Then the reaction was stirred at 0 °C for 6 h. After complete conversion (monitored by TLC), the reaction mixture was concentrated *in vacuo*. Then the solid was washed with petroleum ether .The filtrate were concentrated *in vacuo*. The residue was purified by column chromatography on silica gel (petroleum ether) to get **S-1b** (165 mmol, 91.7%).

n-BuLi (160 mmol, 2.4M in THF) was slowly added to a stirred solution of **S-1b** (80 mmol) in anhydrous THF (13.0 mL) at -78 °C, the reaction mixture was stirred at -78 °C for an additional 4 h. Then the reaction was gradually raised to room temperature for another 4 h. After complete conversion (monitored by TLC), the reaction mixture was extracted with EtOAc. The combined organic layers were concentrated *in vacuo*. The residue was purified by column chromatography on silica gel (petroleum ether) to give **S-1c** (78 mmol, 96%).

Under argon atmosphere, hydrobromic acid was slowly added to **S-1c** (12 mmol) in acetic acid (13 mmol) at 0 °C, the reaction mixture was stirred for 30 min at 0 °C. Then the reaction was quenched by water, and the reaction mixture was extracted with EtOAc. The combined organic layers were concentrated *in vacuo*. The residue was

purified by column chromatography on silica gel (petroleum ether) to get S-1d (8 mmol, 67%).

Pd(OAc)₂ (0.08 mmol) was added to a solution of **S-1d** (2.0 mmol), ArB(OH)₂ (3.0 mmol), K₂CO₃ (6.0 mmol) and 2-dicyclohexylphosphino-2',4',6'-triisopropylbiphenyl (X-Phos, 0.16 mmol) in CH₃CN/H₂O (6.0 mL/3.0 mL). The flask and its contents were put under reduced pressure and then backfilled with argon three times. The reaction mixture was stirred at 80 °C for 24 h under argon atmosphere. After completion, the reaction mixture was cooled to room temperature and extracted with EtOAc, and the combined organic layers were brined, dried over Na₂SO₄ and concentrated *in vacuo*. The residue was dissolved in THF again, *t*-Bu₄N⁺F⁻ (TBAF, 2.0 mmol) was added to the solution of crude product in THF. After complete conversion (monitored by TLC), the reaction mixture was extracted with EtOAc, and the residue was purified by column chromatography on silicagel to give 1 (1.2-1.6 mmol, 60%-80%).

Note: The 1,1-diarylethylenes were unstable at -20-30°C, and easily converted to other products, so the1,1-diarylethylenes should be used in the next reaction ASAP.



4-((triisopropylsilyl)oxy)benzaldehyde (S-1a)

¹**H** NMR (400 MHz, CDCl₃) δ 9.87 (s, 1H), 7.78 (d, J = 8.6 Hz, 2H), 6.97 (d, J = 8.6 Hz, 2H), 1.33-1.24 (m, 3H), 1.10 (d, J = 7.4 Hz, 18H).

¹³C NMR (101 MHz, CDCl₃) δ 190.9, 161.9, 132.0, 130.2, 120.3, 17.9, 12.7.



(4-(2,2-dibromovinyl)phenoxy)triisopropylsilane (S-1b)

¹**H NMR** (400 MHz, CDCl₃) δ 7.45 (d, *J* = 8.0 Hz, 2H), 7.39 (s, 1H), 6.86 (d, *J*= 7.9 Hz, 2H), 1.35-1.20 (m, 3H), 1.10 (t, *J* = 6.0 Hz, 18H).

¹³C NMR (101 MHz, CDCl₃) δ 156.5, 136.4, 129.9, 128.1, 119.8, 87.0, 17.9, 12.7.



(4-ethynylphenoxy)triisopropylsilane (S-1c)

¹**H NMR** (400 MHz, CDCl₃) δ 7.36 (d, J = 8.6 Hz, 2H), 6.81 (d, J = 8.6 Hz, 2H), 2.97 (s, 1H), 1.24 (m, 3H), 1.09 (d, J = 7.4 Hz, 18H).

¹³C NMR (101 MHz, CDCl₃) δ 156.8, 133.6, 120.0, 114.6, 83.8, 75.8, 17.9, 12.7.



(4-(1-bromovinyl)phenoxy)triisopropylsilane (S-1d)

¹**H** NMR (400 MHz, CDCl₃) δ 7.47 (d, J = 8.7 Hz, 2H), 6.83 (d, J = 8.7 Hz, 2H), 6.01 (d, J = 2.0 Hz, 1H), 5.66 (d, J = 2.0 Hz, 1H), 1.34-1.17 (m, 3H), 1.10 (d, J = 7.3 Hz, 18H).

¹³C NMR (101 MHz, CDCl₃) δ 157.1, 131.3, 130.9, 128.6, 119.5, 115.8, 17.9, 12.7.



1-methoxy-4-(1-phenylvinyl)benzene (1aa)

1aa was synthesized according to the procedures.¹



3-(1-(4-methoxyphenyl)vinyl)phenol (1ba)

¹**H NMR** (400 MHz, CDCl₃) δ 7.32 (d, J = 8.7 Hz, 2H), 7.24 (t, J = 7.7 Hz, 1H), 6.97 (d, J = 7.7 Hz, 1H), 6.91 (d, J = 8.7 Hz, 2H), 6.84 (d, J = 7.8 Hz, 2H), 5.71 (s, 1H), 5.41 (d, J = 11.8 Hz, 2H), 3.86 (s, 3H).

¹³**C NMR** (101 MHz, CDCl₃) δ 159.2, 155.4, 149.1, 143.5, 134.0, 129.5, 129.4, 121.0, 115.3, 114.7, 113.6, 113.3, 55.4.

General synthesis of substrate 1ca



According to the procedures with minor revision,² compound **s-1ca** (295 mg, 1.0 mmol) was dissolved in DCM and then cooled to 0 °C. Trifluoroacetic acid was added dropwise, after stirred at rt for 1h, the mixture was concentrated. To the remaining sticky liquid was added 10 mL DCM and washed with saturated aqueous Na₂CO₃, brine and dried over anhydrous Na₂SO₄. After removal of solvent, the residue was purified by flash chromatography on silica gel (petroleum ether/EA = 2:1) to give product **1ca** (150 mg, 77%).



4-(1-phenylvinyl)aniline (1ca)

¹**H** NMR (400 MHz, CDCl₃) δ 7.51 – 7.31 (m, 4H), 7.18 (d, *J* = 8.5 Hz, 2H), 6.67 (d, *J* = 8.5 Hz, 2H), 5.39 (d, *J* = 1.4 Hz, 1H), 5.30 (d, *J* = 1.4 Hz, 1H), 3.74 (s, 2H).

¹³C NMR (101 MHz, CDCl₃) δ 149.7, 146.1, 142.1, 131.7, 129.2, 128.4, 128.0, 127.5, 114.6, 111.8.



4-(1-(p-tolyl)vinyl)phenol (1a)

1a was synthesized according to the procedures.³ **¹H NMR** (400 MHz, CDCl₃) δ 7.36-7.25 (m, 4H), 7.20 (d, *J* = 7.9 Hz, 2H), 6.91-6.83 (m, 2H), 5.39 (dd, *J* = 5.3, 1.3 Hz, 2H),

2.43 (s, 3H).

¹³C NMR (101 MHz, CDCl₃) δ 155.6, 149.4, 139.0, 137.5, 134.2, 129.7, 128.9, 128.3, 115.1, 112.3.



4-(1-phenylvinyl)phenol (1b)

1aa was synthesized according to the procedures.³



4-(1-(4-(tert-butyl)phenyl)vinyl)phenol (1c)

¹**H** NMR (400 MHz, CDCl₃) δ 7.44 (d, *J* = 8.1 Hz, 2H), 7.38 (d, *J* = 7.8 Hz, 2H), 7.33 (d, *J* = 8.0 Hz, 2H), 6.88 (d, *J* = 8.0 Hz, 2H), 5.99 (s, 1H), 5.44 (d, *J* = 5.7 Hz, 2H), 1.43 (s, 10H).

¹³C NMR (101 MHz, CDCl₃) δ 155.2, 150.8, 149.3, 138.8, 134.5, 129.8, 128.0, 125.2, 115.1, 112.6, 34.7, 31.5.



4-(1-(4-chlorophenyl)vinyl)phenol (1d)

1d was synthesized according to the procedures.¹



4-(1-(4-(trifluoromethyl)phenyl)vinyl)phenol (1e)

¹**H** NMR (400 MHz, CDCl₃) δ 7.65 (d, *J* = 8.0 Hz, 2H), 7.50 (d, *J* = 8.0 Hz, 2H), 7.26 (d, *J* = 8.2 Hz, 2H), 6.91 (d, *J* = 8.2 Hz, 2H), 5.54 (s, 1H), 5.46 (s, 1H).

¹³C NMR (101 MHz, CDCl₃) δ 155.6, 148.4, 145.4, 133.4, 129.7, 128.7, 125.2 (q, J = 3.7 Hz), 124.3 (J = 270.0 Hz), 115.4, 114.6.
¹⁹F NMR (376 MHz, CDCl₃) δ -62.3 (s, 3F).

F NMR $(3/0 \text{ MHZ}, \text{CDC}(3) \ 0 \ -02.3 \ (8, \ 3F).$



4-(1-(4-nitrophenyl)vinyl)phenol (1f)

¹**H** NMR (400 MHz, CDCl₃) δ 8.19 (d, *J* = 8.8 Hz, 2H), 7.49 (d, *J* = 8.8 Hz, 2H), 7.17 (d, *J* = 8.6 Hz, 2H), 6.83 (d, *J* = 8.6 Hz, 2H), 5.55 (s, 1H), 5.47 (s, 1H).

¹³C NMR (101 MHz, CDCl₃) δ 155.9, 148.5, 147.8, 147.3, 132.8, 129.6, 129.1, 123.6, 115.9, 115.4.



methyl 4-(1-(4-hydroxyphenyl)vinyl)benzoate (1g)

¹**H NMR** (400 MHz, DMSO) δ 9.61 (d, *J* = 3.7 Hz, 1H),

7.94 (t, *J* = 8.0 Hz, 2H), 7.42 (t, *J* = 8.9 Hz, 2H), 7.09 (t, *J* = 7.3 Hz, 2H), 6.75 (d, *J* = 6.7 Hz, 2H), 5.43 (dd, *J* = 27.7, 9.6

Hz, 2H), 3.85 (d, *J* = 8.7 Hz, 3H).

¹³C NMR (101 MHz, DMSO) δ 166.5, 158.0, 148.6, 146.6, 131.2, 129.7, 129.5, 129.2, 128.7, 115.7, 114.7, 52.6.



4-(1-([1,1'-biphenyl]-4-yl)vinyl)phenol (1h)

¹**H NMR** (400 MHz, CDCl₃) δ 7.61 (d, *J* = 7.5 Hz, 2H), 7.57 (d, *J* = 8.2 Hz, 2H), 7.47-7.32 (m, 5H), 7.23 (t, *J* = 7.8 Hz, 1H), 6.96 (d, *J* = 7.7 Hz, 1H), 6.82 (dd, *J* = 11.4, 2.1 Hz, 2H), 5.49 (d,

J = 13.4 Hz, 2H), 4.89 (s, 1H).

¹³**C NMR** (101 MHz, CDCl₃) δ 155.4, 149.2, 143.2, 140.7, 140.6, 140.2, 129.5, 128.8 (d, *J* = 13.1 Hz), 127.4, 127.0 (d, *J* = 14.6 Hz), 121.0, 115.3, 114.7, 114.5.



4-(1-(4-ethynylphenyl)vinyl)phenol (1i)

¹**H NMR** (400 MHz, CDCl₃) δ 7.51 (d, *J* = 8.0 Hz, 2H), 7.34 (d, *J* = 7.9 Hz, 2H), 7.24 (d, *J* = 8.3 Hz, 2H), 6.87 (d, *J* = 8.3 Hz, 2H), 6.56 (s, 1H), 5.43 (d, *J* = 13.2 Hz, 2H), 3.18 (s, 1H).

¹³C NMR (101 MHz, CDCl₃) δ 155.5, 148.8, 142.3, 133.7, 132.0, 129.7, 128.3, 121.3, 115.3, 113.9, 83.8, 77.9.



4-(1-(3-methoxyphenyl)vinyl)phenol (1j)

¹**H** NMR (400 MHz, CDCl₃) δ 7.34-7.21 (m, 3H), 6.96 (d, *J* = 7.7 Hz, 1H), 6.93-6.86 (m, 2H), 6.82 (t, *J* = 5.7 Hz, 2H), 5.41 (s, 1H), 5.38 (s, 1H), 5.06 (s, 1H), 3.82 (s, 3H).



4-(1-(2-fluorophenyl)vinyl)phenol (1k)

¹**H NMR** (400 MHz, CDCl₃) δ 7.35-7.27 (m, 2H), 7.24-7.18 (m, 2H), 7.14 (td, J = 7.6, 1.2 Hz, 1H), 7.07 (ddd, J = 10.5, 8.1, 0.9 Hz, 1H), 6.84-6.77 (m, 2H), 6.10 (s, 1H), 5.67 (d, J = 1.1 Hz, 1H),

5.31 (t, *J* = 0.9 Hz, 1H).

¹³C NMR (101 MHz, CDCl₃) δ 160.14 (d, J = 248.1 Hz), 155.6, 143.6, 133.2, 131.6 (d, J = 3.7 Hz), 129.57 (d, J = 14.4 Hz), 129.33 (d, J = 8.2 Hz), 128.2, 124.0 (d, J = 3.6 Hz), 115.76 (d, J = 22.3 Hz), 115.3, 115.2. ¹⁹F NMR (375 MHz, CDCl₃) δ -113.4 (s).



4-(1-(thiophen-3-yl)vinyl)phenol (11)

¹**H NMR** (400 MHz, CDCl₃) δ 7.35-7.28 (m, 3H), 7.22-7.14 (m, 2H), 6.85 (d, *J* = 8.7 Hz, 2H), 5.46 (d, *J* = 1.0 Hz, 1H), 5.39 (s, 1H), 5.30 (d, *J* = 1.1 Hz, 1H).

¹³C NMR (100 MHz, CDCl₃) δ 155.4, 143.9, 142.8, 134.1, 129.5, 127.4, 125.4, 123.2, 115.0, 112.3.



4-(1-(4-hydroxyphenyl)vinyl)benzaldehyde (1q)

1q was obtained by the same procedure for the compound 1c ¹H NMR (400 MHz, CDCl₃) δ 10.02 (s, 1H), 7.86 (d, *J* = 8.2 Hz, 2H), 7.52 (d, *J* = 8.2 Hz, 2H), 7.18 (d, *J* = 8.6 Hz, 2H),

6.86 (d, *J* = 8.4 Hz, 2H), 5.51 (s, 1H), 5.45 (s, 1H).

¹³C NMR (101 MHz, CDCl₃) δ 192.1, 155.8, 148.6, 148.1, 135.6, 133.1, 129.7, 129.6, 128.9, 115.3, 115.1.

General synthesis of substrates 1n-1p



1n-1p were synthesized according to the procedures previously reported.³



2-fluoro-4-(1-(p-tolyl)vinyl)phenol (1n)

¹**H NMR** (400 MHz, CDCl₃) δ 7.21 (d, *J* = 8.0 Hz, 2H), 7.14 (d, *J* = 7.6 Hz, 2H), 7.07 (dd, *J* = 11.6, 2.0 Hz, 1H), 7.03 (ddd, *J* = 11.6, 2.4, 1.0 Hz, 1H), 6.94 (t, *J* = 8.8 Hz, 1H), 5.35 (s, 1H),

5.30 (brs, 1H), 2.37 (s, 3H). ¹³C NMR (101 MHz, CDCl₃) δ 150.6 (d, J = 237.0 Hz), 138.25, 137.69, 148.5 (d, J = 1.8 Hz), 143.1 (d, J = 14.5 Hz), 134.9 (d, J = 6.0 Hz), 128.90, 128.15, 124.6 (d, J = 3.2 Hz), 116.7 (d, J = 2.1 Hz), 115.3 (d, J = 18.6 Hz), 113.16, 21.14. ¹⁹F NMR (376 MHz, CDCl₃) δ -141.2 (s, 1F).



2-methyl-4-(1-(p-tolyl)vinyl)phenol (10)

¹**H NMR** (400 MHz, CDCl₃) δ 7.30 (s, 2H), 7.20 (s, 3H), 7.11 (s, 1H), 6.77 (d, *J* = 8.2 Hz, 1H), 5.38 (d, *J* = 2.9 Hz, 2H), 5.13 (s, 1H), 2.43 (s, 3H), 2.29 (s, 3H).

¹³C NMR (101 MHz, CDCl₃) δ 153.6, 149.5, 139.1, 137.5, 134.4, 131.0, 128.9, 128.3, 127.2, 123.4, 114.6, 112.3, 21.2, 15.9.



3-fluoro-4-(1-(*p*-tolyl)vinyl)phenol (1p)

¹**H** NMR (400 MHz, CDCl₃) δ 7.24 (d, *J* = 8.1 Hz, 2H), 7.15 (t, *J* = 7.9 Hz, 3H), 6.68-6.56 (m, 2H), 5.66 (s, 1H), 5.34 (s, 1H), 5.17 (s, 1H), 2.37 (s, 3H).

¹³C NMR (126 MHz, CDCl₃) δ 160.7 (d, J = 248.9 Hz), 156.4 (d, J = 11.4 Hz), 143.7, 138.1, 137.6, 132.1 (d, J = 5.5 Hz), 129.0, 126.8, 122.0 (d, J = 14.5 Hz), 115.6 (d, J = 2.2 Hz), 111.0 (d, J = 3.1 Hz), 103.4 (d, J = 25.8 Hz), 21.2. ¹⁹F NMR (376 MHz, CDCl₃) δ -111.0 (s, 1F).

Substituted indoles were used in the Cu/CPA-catalyzed asymmetric three-component radical-initiated 1,2-dicarbofunctionalization of alkenes



Synthesis of the catalyst (S)-A5

(S)-6,6'-bis(2,4,6-tricyclohexylphenyl)- 4,4'-dimethyl- 2,2',3,3'-tetrahydro-1,1' -spirobi[indene]-7,7'-diol (S-1f)



(2,4,6-Tricyclohexylphenyl)magnesium bromide. Mg (320 mg, 13 mmol), (2-bromobenzene-1,3,5-triyl)tricyclohexane (4.02 g, 10 mmol) and anhydrous THF (30.0 mL) were added to an oven-dried resealable Schlenk tube (100 mL) under argon atmosphere, then one bean of I₂ was added. The reaction mixture was heated to reflux, and then Me₃SiCl (150 μ L) and 1,2-dibromoethane (150 μ L) were added. The resultant solution was heated at 80 °C for 2 h, until only trace amounts of magnesium metal remained unreacted. The resulting Grignard solution was used directly in the next step.

(S)-6,6'-diiodo-4,4'-dimethyl-2,2',3,3'-tetrahydro-1,1'-spirobi[indene]-7,7'-diol S-1e (1.33g, 2.5 mmol) and anhydrous THF (5 mL) were added into an oven-dried resealable Schlenk tube (100 ml) under argon atmosphere. The resulting solution was placed in an ice bath, and NaH (380 mg) was slowly added under argon atmosphere. Then the resulting mixture was warmed up to room temperature and stirred for 15 minutes. Pd(OAc)₂ (100 mg, 0.44 mmol) was added, followed by the Grignard reagent prepared before, and the reaction mixture was heated to 80 °C for 24 h. Upon cooling to room temperature, H₂O was carefully added to quench the residual Grignard reagent and sodium hydride. HCl (2 M, 20 mL) and EtOAc (30 mL) were added and the mixture stirred for 5 minutes, then the resulting mixture was filtered through a pad of celite. The resulting phases were separated, and the aqueous phase was extracted with EtOAc (3 times). The combined organic layers were washed with brine, dried over anhydrous Na₂SO₄, filtered and concentrated in vacuo. The resulting residue was purified by flash column chromatography (silica gel, eluent: *n*-hexane/ethyl acetate = 100/0-25/1) to give the S-1f (1.54g, 1.67 mmol, 67%) as a white solid.



(*S*)-6,6'-Bis(2,4,6-tricyclohexylphenyl)- 4,4'-dimethyl -2,2',3,3'-tetrahydro-1,1'-spirobi[indene]-7,7'-diol (S-1f) ¹H NMR (500 MHz, CDCl₃) δ 6.98 (s, 2H), 6.95 (s, 2H), 6.65 (s, 2H), 4.11 (d, *J* = 1.6 Hz, 2H), 2.97 (dd, *J* = 9.5, 3.7 Hz, 4H), 2.49 (t, *J* = 9.5 Hz, 2H), 2.40-2.31 (m, 2H), 2.30-2.17 (m, 10H), 2.11 (t, *J* = 11.7 Hz, 2H), 1.87 (s, 8H), 1.82-1.58 (m, 14H), 1.51-0.82 (m, 38H). ¹³C NMR (126 MHz, CDCl₃) δ 147.9, 147.4, 147.3, 147.1, 142.6, 132.1, 131.0, 130.2, 124.5, 124.3, 122.1, 121.9, 59.2, 44.9, 41.3, 40.8, 37.3, 35.0, 34.7, 34.6, 34.42, 34.38, 34.1, 29.8, 27.1, 27.0, 27.0, 26.7, 26.3, 26.13, 26.08, 18.5.

HRMS (ESI) m/z calcd. for C₆₇H₈₈NaO₂ [M+Na]⁺ 947.66765, found 947.66833.



Chiral Phosphoric Acid (S)-A5. S-1f (925 mg, 1 mmol) was suspended in 15 mL of pyridine and treated with 1 mL of freshly distilled POCl₃. The resulting solution was stirred at 100 °C for 6 days. After cooling to room temperature, H₂O (3 mL) was added carefully and the resulting mixture was heated to 100 °C for 2 days. The reaction mixture was acidified with HCl (2N) and extracted with EtOAc for three times. The combined organic layers were washed with 2N HCl for two times, dried over anhydrous Na₂SO₄, filtered and concentrated *in vacuo*. The resulting residue was purified by flash column chromatography (silica gel, eluted with *n*-hexane /DCM=10/1 then *n*-hexane/ EtOAc=4/1) to give the product as a slight yellow solid in 56% yield (552 mg, 0.56 mmol) and 200 mg of substrate was recovered.



(S)-A5

¹**H** NMR (500 MHz, CDCl₃) δ 7.00 (d, J = 1.3 Hz, 2H), 6.94 (d, J = 1.3 Hz, 2H), 6.87 (s, 2H), 3.07-2.96 (m, 2H), 2.90 (dd, J = 16.1, 8.0 Hz, 2H), 2.77 (s, 9H), 2.54-2.41 (m, 4H), 2.39-2.22 (m, 10H), 2.14 (dd, J = 19.5, 10.8 Hz, 2H), 2.01-1.81 (m, 12H), 1.74 (t, J = 14.3 Hz, 7H), 1.62 (t, J = 11.3 Hz, 8H), 1.53-0.87 (m, 32H).

¹³C NMR (126 MHz, CDCl₃) δ 147.1, 146.7, 146.4, 142.8, 142.8, 141.4, 141.3, 138.9, 138.9, 132.9, 132.7, 131.2, 131.1, 130.2, 122.5, 121.5, 60.7, 44.7, 41.7, 41.5, 38.1, 37.6, 35.5, 34.50, 34.48, 33.4, 33.0, 28.8, 27.6, 27.5, 27.0, 26.8, 26.6, 26.4, 26.2, 18.4.
³¹P NMR (162 MHz, CDCl₃) δ -9.51(s, 1P).

HRMS (ESI) m/z calcd. for C₆₇H₈₈PO₄ [M+H]⁺ 987.64147, found 987.64404.

General procedure for initial exploration and screening of reaction conditions

Cul (10 mol%) n-C₄F₉ (S)-A1 (10 mol%) n-C₄F₉SO₂Cl (3a) Ag₂CO₃ (0.6 eq) DCM, 25 °C, 48 h 5 6 (S)-A1: Ar = 4-Ph-C₆H, \mathbb{R}^2 \mathbb{R}^1 4 y (%) ee (%) y (%) 1 5 y (%) Entry 6 4Aa 20 7 5Aa 53 1 1aa 4-H OMe 6Aa 24 2 1ba 3-OH OMe 4Ba 54 43 5Ba 44 6Ba 0 3 1a **4-OH** Me **4**A 10 68 5Ca 44 6Ca 43 4 1ca 4-NH2 Η 4Ca 0 5Da 68 6Da 0

Table 1. Initial Exploration of Reaction Conditions^a

^{*a*}Reaction conditions: **1** (0.1 mmol), **2a** (0.12 mmol), n-C₄F₉SO₂Cl (0.12 mmol), CuI (10 mol%), Ag₂CO₃ (0.06 mmol), CPA (10 mol%), DCM (1.0 mL) under argon. ^{*b*} Isolated yield. ^{*c*}Ee value on HPLC.

Under argon atmosphere, an oven-dried resealable Schlenk tube equipped with a magnetic stir bar was charged with 1,1-diarylalkene substrate **1** (0.1 mmol, 1.0 equiv.), 5-bromo-1*H*-indole (0.12 mmol, 1.2 equiv.), CuI (1.9 mg, 0.01 mmol, 10 mol%), chiral phosphoric acid ((*S*)-**A1** (6.18 mg, 0.01 mmol, 10 mol%), *n*-C4F9SO₂Cl (**3a**) (38.2 mg, 0.12 mmol, 1.2 equiv.), Ag₂CO₃ (16.56 mg, 0.06 mmol, 0.6 equiv.), and anhydrous DCM (1.0 mL) at 25 °C, then the sealed tube was stirred at 25 °C for 48 h. Upon completion (monitored by TLC), the reaction mixture was directly purified by a silica gel chromatography [eluent: *n*-hexane/ethyl acetate = 20/0-2/1, using *n*-hexane (100%) to remove the solvent (DCM) at first] to afford the desired product **4**, **5** and **6**. *Note: Since the reaction is sensitive to water and air, Schlenk tube and the reagents must be dried prior to use.*



The racemate was prepared following the same procedure described above using CuI (3.0 mg, 0.015 mmol, 15 mol%) and *rac*-A-NHTf (6.3 mg, 0.01 mmol, 10 mol%) as catalyst at 0 °C in anhydrous DCM (1.0 mL) for 48 h. Upon completion (monitored by TLC), the solvent was removed *in vacuo*, and the residue was purified by a silica gel column chromatography (eluent: *n*-hexane/ethyl acetate = 20/0-2/1) to give the desired product.

Note: Since the reaction is sensitive to water and air, Schlenk tube and the reagents must be dried prior to use.



(*R*)-5-bromo-3-(3,3,4,4,5,5,6,6,6-nonafluoro-1-(4-methoxyphe nyl)-1-phenylhexyl)-1*H*-indole (4Aa)

The reaction was conducted on a 0.1 mmol scale according to the above general procedure. The product was purified by silica gel flash column chromatography (n-hexane/ethyl acetate = 100:0 to

3:1) to afford desired product **4Aa** (12.5 mg, 20 %) as sticky light yellow oil. **HPLC** analysis: Chiralcel IA (hexane/*i*-PrOH = 85/15, flow rate 0.5 mL/min, λ = 254 nm), $t_{\rm R}$ (minor) = 14.66 min, $t_{\rm R}$ (major) = 15.89 min.

¹**H NMR** (500 MHz, CDCl₃) δ 8.18 (s, 1H), 7.40 (d, J = 7.6 Hz, 2H), 7.33-7.27 (m, 4H), 7.27-7.20 (m, 3H), 7.07 (d, J = 1.5 Hz, 2H), 6.86-6.79 (m, 2H), 3.81 (s, 3H), 3.53 (t, J = 18.6 Hz, 2H).

¹³**C NMR** (126 MHz, CDCl₃) δ 158.0, 145.0, 136.5, 135.4, 129.4, 128.2, 128.1, 128.0, 126.5, 125.5, 124.9, 124.2, 120.4, 113.3, 112.7, 112.7, 117.8-106.1 (m), 55.2, 49.1, 39.1 (t, *J* = 19.5 Hz).

¹⁹**F** NMR (376 MHz, CDCl₃) δ -80.5 ~ -81.4 (m, 3F), -108.6 (t, *J* = 12.1 Hz, 2F), -124.2 (d, *J* = 5.8 Hz, 2F), -125.2 ~ -126.0 (m, 2F).

HRMS (ESI) m/z calcd. for C₂₇H₁₈NOBrF₉ [M-H]⁻ 622.04335, found 622.04303.



(*E*/*Z*)-1-methoxy-4-(3,3,4,4,5,5,6,6,6-nonafluoro-1-phenylhex-1en-1-yl)benzene (5Aa)

The product was purified by silica gel flash column chromatography (*n*-hexane/ethyl acetate = 100:0 to 10:1) to afford desired product **5Aa** (22.7 mg, 53 %) as sticky colorless oil.

¹**H** NMR (500 MHz, CDCl₃, observed as a mixture of Z and *E*-isomers) δ 7.49-7.32 (m, 3H, *E*+*Z*), 7.27-7.24 (m, 2H, *E*+*Z*), 7.20 (t, *J* = 7.5 Hz, 2H, *E*+*Z*), 6.94 (d, *J* = 8.7 Hz, 2H, *Z*), 6.88 (d, *J* = 8.8 Hz, 2H, *E*), 6.05 (t, *J* = 14.8 Hz, 1H, *E*+*Z*), 3.87 (s, 3H, *Z*), 3.84 (s, 3H, *E*).

¹³C NMR (126 MHz, CDCl₃, observed as a mixture of Z and *E*-isomers) δ 160.8 (*E*), 159.8 (*Z*), 154.4 (*Z*), 153.8 (*E*), 153.7, 153.7, 141.4, 137.7, 133.0, 130.6, 129.8, 129.5, 129.4, 129.04, 129.02, 128.4, 128.23, 128.15, 127.8, 118.9-105.8 (m), 113.8, 113.3, 110.5 (t, *J* = 21.1 Hz), 55.4 (*E*), 55.2 (*Z*).

¹⁹**F NMR** (376 MHz, CDCl₃, observed as a mixture of Z and *E*-isomers) δ -81.0 ~ -81.1 (m, 3F, *E*+*Z*), -103.0 ~ -103.3 (m, 2F, *E*), -103.3 ~ -103.4 (m, 2F, *Z*), -123.8 ~ -124.1 (m, 2F, *E*+*Z*), -125.5 ~ -125.8 (m, 2F, *E*+*Z*).



5-bromo-3-(1-(4-methoxyphenyl)-1-phenylethyl)-1H-indole (6Aa)

The product was purified by silica gel flash column chromatography (*n*-hexane/ethyl acetate = 100:0 to 3:1) to afford desired product **6Aa** (9.7 mg, 24 %) as a light yellow solid.

¹**H NMR** (500 MHz, CDCl₃) δ 7.98 (s, 1H), 7.35-7.20 (m, 8H), 7.14 (d, *J* = 8.8 Hz, 2H), 6.83 (d, *J* = 8.8 Hz, 2H), 6.47 (d, *J* = 2.3 Hz, 1H), 3.82 (s, 3H), 2.24 (s, 3H).

¹³C NMR (126 MHz, CDCl₃) δ 157.7, 148.4, 140.3, 135.8, 129.3, 128.2, 127.9, 126.0, 125.6, 124.9, 124.7, 124.4, 113.2, 112.6, 112.5, 55.2, 47.3, 29.4. HRMS (ESI) m/z calcd. for C₂₃H₁₉NOBr [M-H]⁻ 404.06555, found 404.06516.



(*R*)-3-(1-(5-bromo-1*H*-indol-3-yl)-3,3,4,4,5,5,6,6,6-nonafluor o-1-(4-methoxyphenyl)hexyl)phenol (4Ba)

The reaction was conducted on a 0.1 mmol scale according to the above general procedure. The product was purified by silica gel flash column chromatography (n-hexane/ethyl acetate =

100:0 to 3:1) to afford desired product **4Ba** (34.5 mg, 54 %) as sticky light yellow oil. **HPLC** analysis: Chiralcel IA (*n*-hexane/*i*-PrOH = 85/15, flow rate 0.4 mL/min, λ = 240 nm), $t_{\rm R}$ (minor) = 16.43 min, $t_{\rm R}$ (major) = 19.00 min.

¹**H NMR** (500 MHz, CDCl₃) δ 8.18 (s, 1H), 7.34-7.26 (m, 2H), 7.22 (s, 2H), 7.16 (t, *J* = 8.0 Hz, 1H), 7.10 (s, 1H), 7.04 (d, *J* = 1.6 Hz, 1H), 6.98 (d, *J* = 7.9 Hz, 1H), 6.86 (d, *J* = 1.9 Hz, 1H), 6.82 (d, *J* = 8.9 Hz, 2H), 6.69 (dd, *J* = 8.0, 2.2 Hz, 1H), 5.20 (s, 1H), 3.80 (s, 3H), 3.50 (t, *J* = 18.6 Hz, 2H).

¹³C NMR (126 MHz, CDCl₃) δ 158.0, 155.2, 147.2, 136.2, 135.4, 129.4, 129.2, 128.2, 125.6, 124.9, 124.2, 120.8, 120.3, 120.0, 119.4-106.8 (m) 115.4, 113.6, 112.7, 112.6, 55.2, 49.0, 39.0 (t, *J* = 19.4 Hz).

¹⁹**F NMR** (376 MHz, CDCl₃) δ -81.0 (t, J = 9.7 Hz, 3F), -108.7 (t, J = 13.8 Hz, 2F), -123.9 ~ -124.5 (m, 2F), -125.5 ~ -125.9 (m, 2F).

HRMS (ESI) m/z calcd. for $C_{27}H_{18}F_9BrNO_2$ [M-H]⁻ 638.03827, found 638.03754.



(*E/Z*)-3-(3,3,4,4,5,5,6,6,6-nonafluoro-1-(4-methoxyphenyl)h ex-1-en-1-yl)phenol (5Ba)

The product was purified by silica gel flash column chromatography (*n*-hexane/ethyl acetate = 100:0 to 10:1) to afford desired product **5Ba** (19.5 mg, 44 %) as sticky colorless

oil.

¹**H NMR** (500 MHz, CDCl₃) δ 7.36-7.09 (m, 3H, *E*+*Z*), 6.98-6.77 (m, 4H, *E*+*Z*), 6.70 (d, *J* = 10.2 Hz, 1H, *E*+*Z*), 6.02 (td, *J* = 14.7, 8.3 Hz, 1H, *E*+*Z*), 5.14 (s, 1H, *E*+*Z*), 3.85 (d, *J* = 15.4 Hz, 3H, *E*+*Z*).

¹³C NMR (126 MHz, CDCl₃, observed as a mixture of *Z* and *E*-isomers) δ 160.8 (*E*), 159.8 (*Z*), 155.6 (*Z*), 155.1 (*E*), 153.90 (*Z*), 153.2 (*E*), 143.0, 139.2 (*E*), 132.7, 130.6, 129.7, 129.6, 129.3, 129.2, 121.7, 120.6, 116.8, 116.4, 116.0, 115.3, 115.2, 113.9, 113.3, 112.4 (t, *J* = 20.7 Hz), 110.5 (t, *J* = 21.2 Hz), 55.4 (*E*), 55.3 (*Z*).

¹⁹**F NMR** (376 MHz, CDCl₃, observed as a mixture of Z and *E*-isomers) δ -80.5 ~ -81.3 (m, 3F, *E*+*Z*), -103.46 (t, *J* = 11.1 Hz, 2F, *E*+*Z*), -123.7 ~ -124.3 (m, 2F, *E*+*Z*), -125.4 ~ -125.8 (m, 2F, *E*+*Z*).

HRMS (ESI) m/z calcd. for $C_{19}H_{12}F_9O_2$ [M-H]⁻ 443.06991, found 443.06921.



(*S*)-4-(1-(5-bromo-1*H*-indol-3-yl)-3,3,4,4,5,5,6,6,6-nonafluor o-1-(*p*-tolyl)hexyl)phenol (4A)

The reaction was conducted on a 0.1 mmol scale according to

the above general procedure. The product was purified by silica gel flash column chromatography (*n*-hexane/ethyl acetate = 100:0 to 3:1) to afford desired product **4A** (6.2 mg, 10 %) as sticky light yellow oil.

HPLC analysis: Chiralcel IA (*n*-hexane/*i*-PrOH = 85/15, flow rate 0.5 mL/min, λ = 240 nm), t_R (major) = 10.57 min, t_R (minor) = 11.72 min.

¹**H** NMR (500 MHz, CDCl₃) δ 8.13 (s, 1H), 7.26-7.20 (m, 6H), 7.07 (d, *J* = 7.5 Hz, 3H), 7.04 (d, *J* = 2.4 Hz, 1H), 6.79-6.71 (m, 2H), 4.90 (br s, 1H), 3.57-3.37 (m, 2H), 2.30 (s, 3H).

¹³**C NMR** (126 MHz, CDCl₃) δ 154.0, 142.0, 136.9, 136.1, 135.4, 129.6, 128.8, 128.2, 127.9, 125.4, 124.8, 124.2, 120.6, 114.8, 112.7, 112.6, 119.9-105.6 (m), 48.8, 39.2 (t, *J* = 19.6 Hz), 20.9.

¹⁹F NMR (376 MHz, CDCl₃) δ -80.7 ~ -81.4 (m, 3F), -108.4 ~ -109.0 (m, 2F), -124.2 (d, J = 8.2 Hz, 2F), -125.1 ~ -126.2 (m, 2F).

HRMS (ESI) m/z calcd. for $C_{27}H_{18}ONBrF_9 [M-H]^- 622.0434$, found 622.0455.



(*E*)-4-(3,3,4,4,5,5,6,6,6-nonafluoro-1-(4-methoxyphenyl)hex -1-en-1-yl)phenol (5Ca)

Ho 5Ca Me The product was purified by silica gel flash column chromatography (*n*-hexane/ethyl acetate = 100:0 to 10:1) to afford desired product **5Ca** (18.8 mg, 44 %) as sticky colorless oil.

¹**H NMR** (500 MHz, CDCl₃) δ 7.20 (d, *J* = 7.8 Hz, 2H), 7.14 (dd, *J* = 12.5, 8.2 Hz, 4H), 6.80 (d, *J* = 8.6 Hz, 2H), 5.99 (t, *J* = 14.8 Hz, 1H), 5.22 (s, 1H), 2.42 (s, 3H).

¹³**C NMR** (126 MHz, CDCl₃) δ 156.8, 153.9, 138.2, 134.7, 133.7, 129.7, 129.0, 128.5, 115.3, 119.0-106.5 (m), 110.4 (t, *J* = 21.0 Hz), 21.3.

¹⁹**F NMR** (376 MHz, CDCl₃) δ -81.0 (s, 3F), -103.1 (s, 2F), -123.9 (s, 2F), -125.6 (s, 2F).

HRMS (ESI) m/z calcd. for $C_{19}H_{12}OF_9 [M-H]^- 427.07499$, found 427.07535.



4-(1-(5-bromo-1*H*-indol-3-yl)-1-(*p*-tolyl)ethyl)phenol (6Ca)

The product was purified by silica gel flash column chromatography (*n*-hexane/ethyl acetate = 100:0 to 3:1) to afford desired product **6Ca** (17.4 mg, 43 %) as a light yellow solid.

¹**H** NMR (500 MHz, CDCl₃) δ 7.93 (s, 1H), 7.24 (s, 1H), 7.20 (d, J = 1.0 Hz, 2H), 7.09-7.04 (m, 5H), 7.03 (d, J = 2.1 Hz, 1H), 6.74-6.67 (m, 2H), 6.43 (d, J = 2.5 Hz, 1H), 4.83 (s, 1H), 2.31 (s, 3H), 2.16 (s, 3H).

¹³**C NMR** (126 MHz, CDCl₃) δ 153.6, 145.4, 140.7, 135.8, 135.5, 129.5, 128.7, 128.1, 125.7, 124.8, 124.6, 124.4, 114.7, 112.6, 112.4, 46.9, 29.5, 21.0.

HRMS (ESI) m/z calcd. for C₂₃H₁₉ONBr [M-H]⁻ 404.0656, found 404.0667.



(*E*)-4-(3,3,4,4,5,5,6,6,6-nonafluoro-1-phenylhex-1-en-1-yl)anili ne (5Da)

The product was purified by silica gel flash column chromatography (*n*-hexane/ethyl acetate = 100:0 to 10:1) to afford

desired product 5Da (28.0 mg, 68 %) as light yellow oil. Note: 5Da is unstable at rt

under air atmosphere, easily converted to other products.

¹**H NMR** (500 MHz, CDCl₃) δ 7.41-7.34 (m, 3H), 7.22 (dd, *J* = 6.4, 2.9 Hz, 2H), 7.04 (d, *J* = 8.6 Hz, 2H), 6.63-6.55 (m, 2H), 5.97 (t, *J* = 15.0 Hz, 1H), 3.80 (br s, 2H). ¹³**C NMR** (126 MHz, CDCl₃) δ 153.9, 147.9, 137.9, 130.5, 129.3, 129.1, 128.0, 127.7,

114.5, 125.3-110.8 (m), 108.9 (t, J = 21.0 Hz).

¹⁹**F NMR** (376 MHz, CDCl₃) δ -79.4 ~ -83.5 (m, 3F), -102.2 ~ -103.0 (m, 2F), -123.3 ~ -124.4 (m, 2F), -125.3 ~ -126.0 (m, 2F).

HRMS (ESI) m/z calcd. for C₁₈H₁₃NF₉ [M+H]⁻ 414.08988, found 414.08859.

Scheme S1. Asymmetric Intermolecular Three-Component Radical-Initiated Dicarbofunctionalization of *meta*-Phenol Substrate 1ba



The reaction was conducted on a 0.025 mmol scale according to the above general procedure in the presence of (S)-A8. The product was purified by silica gel flash column chromatography (*n*-hexane/ethyl acetate = 100:0 to 3:1) to afford desired product **4Ba** (5.6 mg, 35 %, 53% ee).

General procedure A: Cu/CPA-catalyzed asymmetric three-component radical -initiated perfluoroalkylarylation or difluoroacetylarylation of 1,1-diarylalkene



Under argon, an oven-dried resealable Schlenk tube equipped with a magnetic stir bar was charged with 1,1-diarylalkene substrate 1 (0.1 mmol, 1.0 equiv.), substituted indole (0.12 mmol, 1.2 equiv.), CuI (1.9 mg, 0.01 mmol, 10 mol%), chiral phosphoric acid ((*S*)-A5 (9.86 mg, 0.01 mmol, 10 mol%), *n*-C4F9SO₂Cl (**3a**, 38.2 mg, 0.12 mmol, 1.2 equiv.), Ag₂CO₃ (16.56 mg, 0.06 mmol, 0.6 equiv.), and anhydrous DCM (1.0 mL) at 0 °C, then the sealed tube was stirred at 0 °C for 40-96 h. Upon completion (monitored by TLC), the reaction mixture was directly purified by a silica gel chromatography [eluent: *n*-hexane/ethyl acetate = 20/0-2/1, using *n*-hexane (100%) to remove the solvent (DCM) at first] to afford the desired product **4**.

The difluoroacetyl-containing product **5A** was prepared following the same procedure described above conducting on 0.025 mmol scale with 1,1-diarylalkene substrate **1** (5.25 mg, 0.025 mmol, 1.0 equiv.), MeO₂CCF₂SO₂Cl (**3b**, 6.4 mg, 0.03 mmol, 1.2 equiv.), Ag₂CO₃ (3.5 mg, 0.025 mmol, 0.5 equiv.), (*R*)-3,3'-(3,5-(Ph)₂C₆H₃)₂-8*H* -BINOL-derived CPA (2.0 mg, 0.0025 mmol, 10 mol%) at -30 °C for 72 h. *Note: Since the reaction is sensitive to water and air, Schlenk tube and the reagents must be dried prior to use.*



The racemate was prepared following the same procedure described above using CuI (3.0 mg, 0.015 mmol, 15 mol%) and *rac*-A-NHTf (6.3 mg, 0.01 mmol, 10 mol%) as catalyst at 0 °C in anhydrous DCM (1.0 mL) for 48-96 h. Upon completion (monitored by TLC), the solvent was removed *in vacuo*, and the residue was purified by a silica gel column chromatography (eluent: *n*-hexane/ethyl acetate = 20/0-2/1) to give the desired product.

Note: Since the reaction is sensitive to water and air, Schlenk tube and the reagents must be dried prior to use.



(*S*)-4-(1-(5-bromo-1*H*-indol-3-yl)-3,3,4,4,5,5,6,6,6-nonafluor o-1-(*p*-tolyl)hexyl)phenol (4A)

The reaction was conducted on a 0.1 mmol scale according to the above general procedure A. The product was purified by silica gel flash column chromatography (*n*-hexane/ethyl acetate

= 100/0 to 2/1) to afford desired product **4A** (51.1 mg, 82 %) as sticky light yellow oil. **HPLC** analysis: Chiralcel IA (*n*-hexane/*i*-PrOH = 85/15, flow rate 0.5 mL/min, λ = 240 nm), $t_{\rm R}$ (minor) = 10.39 min, $t_{\rm R}$ (major) = 11.52 min.

¹**H** NMR (500 MHz, CDCl₃) δ 8.13 (s, 1H), 7.26-7.20 (m, 6H), 7.07 (d, J = 7.5 Hz, 3H), 7.04 (d, J = 2.4 Hz, 1H), 6.79-6.71 (m, 2H), 4.90 (br s, 1H), 3.57-3.37 (m, 2H), 2.30 (s, 3H).

¹³C NMR (126 MHz, CDCl₃) δ 154.0, 142.0, 136.9, 136.1, 135.4, 129.6, 128.8, 128.2, 127.9, 125.4, 124.8, 124.2, 120.6, 114.8, 112.7, 112.6, 119.9-105.6 (m), 48.8, 39.2 (t, *J* = 19.6 Hz), 20.9.

¹⁹**F NMR** (376 MHz, CDCl₃) δ -80.7 ~ -81.4 (m, 3F), -108.4 ~ -109.0 (m, 2F), -124.2 (d, J = 8.2 Hz, 2F), -125.1 ~ -126.2 (m, 2F).

HRMS (ESI) m/z calcd. for C₂₇H₁₈ONBrF₉[M-H]⁻ 622.0434, found 622.0455.



(*S*)-4-(1-(5-bromo-1*H*-indol-3-yl)-3,3,4,4,5,5,6,6,6-nonafluor o-1-phenylhexyl)phenol (4B)

The reaction was conducted on a 0.1 mmol scale according to the above general procedure A. The product was purified by silica gel flash column chromatography (n-hexane/ethyl acetate

= 100/0 to 2/1) to afford desired product **4B** (41.2 mg, 66 %) as sticky light yellow oil. **HPLC** analysis: Chiralcel AD3 (*n*-hexane/*i*-PrOH = 85/15, flow rate 0.3 mL/min, λ = 230 nm), *t*_R (minor) = 17.21 min, *t*_R (major) = 21.59 min.

¹**H NMR** (500 MHz, CDCl₃) δ 8.18 (s, 1H), 7.40 (d, *J* = 7.8 Hz, 2H), 7.35-7.18 (m, 7H), 7.07 (s, 2H), 6.81-6.71 (m, 2H), 5.01 (br s, 1H), 3.62-3.42 (m, 2H).

¹³**C NMR** (126 MHz, CDCl₃) δ 154.0, 144.9, 136.7, 135.4, 129.6, 128.2, 128.1, 128.0, 126.6, 125.4, 124.9, 124.2, 120.4, 114.8, 112.72, 112.66, 120.8-108.9 (m), 49.1, 39.2 (t, *J* = 19.5 Hz).

¹⁹**F NMR** (376 MHz, CDCl₃) δ -81.0 (t, J = 9.7 Hz, 3F), -108.6 (s, 2F), -124.2 (d, J = 4.3 Hz, 2F), -125.0 ~ -126.3 (m, 2F).

HRMS (ESI) m/z calcd. for C₂₆H₁₆ONBrF₉[M-H]⁻ 608.0277, found 608.0290.



(*S*)-4-(1-(5-bromo-1*H*-indol-3-yl)-1-(4-(*tert*-butyl)phenyl)-3,3 ,4,4,5,5,6,6,6-nonafluorohexyl)phenol (4C)

The reaction was conducted on a 0.1 mmol scale according to the above general procedure A. The product was purified by silica gel flash column chromatography (n-hexane/ethyl acetate

= 100/0 to 2/1) to afford desired product 4C (56.5 mg, 85 %) as sticky light yellow oil. HPLC analysis: Chiralcel IA (*n*-hexane/*i*-PrOH = 90/10, flow rate 0.2 mL/min, λ = 240 nm), $t_{\rm R}$ (minor) = 34.07 min, $t_{\rm R}$ (major) = 35.93 min. ¹**H** NMR (500 MHz, CDCl₃) δ 8.12 (s, 1H), 7.35-7.24 (m, 6H), 7.21 (q, J = 8.6 Hz, 2H), 7.05 (d, J = 11.6 Hz, 2H), 6.75 (d, J = 8.6 Hz, 2H), 3.69-3.37 (m, 2H), 1.32 (s, 9H).

¹³**C NMR** (126 MHz, CDCl₃) δ 154.0, 149.3, 142.2, 136.9, 135.4, 129.7, 128.3, 127.5, 125.6, 125.0, 124.7, 124.3, 120.5, 114.8, 112.7, 112.6, 120.8-106.5 (m), 48.7, 39.1 (t, *J* = 19.4 Hz) 34.4, 31.3.

¹⁹F NMR (376 MHz, CDCl₃) δ -81.0 (t, J = 9.6 Hz, 3F), -107.5 ~ -109.9 (m, 2F), -123.8 ~ -124.7 (m, 2F), -125.6 (t, J = 11.6 Hz, 2F).

HRMS (ESI) m/z calcd. for C₃₀H₂₄ONBrF₉ [M-H]⁻ 664.0903, found 664.0918.



(*S*)-4-(1-(5-bromo-1*H*-indol-3-yl)-1-(4-chlorophenyl)-3,3,4,4 ,5,5,6,6,6-nonafluorohexyl)phenol (4D)

The reaction was conducted on a 0.1 mmol scale according to the above general procedure A. The product was purified by silica gel flash column chromatography (n-hexane/ethyl acetate

= 100/0 to 2/1) to afford desired product **4D** (36.1 mg, 56 %) as sticky light yellow oil. **HPLC** analysis: Chiralcel IA (*n*-hexane/*i*-PrOH = 85/15, flow rate 0.3 mL/min, λ = 230 nm), $t_{\rm R}$ (minor) = 19.09 min, $t_{\rm R}$ (major) = 22.02 min.

¹**H** NMR (500 MHz, CDCl₃) δ 8.20 (s, 1H), 7.33 (d, J = 8.6 Hz, 2H), 7.30-7.24 (m, 4H), 7.22 (d, J = 8.7 Hz, 2H), 7.12-7.01 (m, 2H), 6.76 (d, J = 8.7 Hz, 2H), 4.96 (s, 1H), 3.48 (t, J = 18.4 Hz, 2H).

¹³C NMR (126 MHz, CDCl₃) δ 154.2, 143.2, 136.4, 135.4, 132.4, 129.6, 129.5, 128.2, 127.9, 125.3, 125.1, 124.0, 120.0, 115.0, 112.8, 120.2-107.8 (m), 48.8, 39.2 (t, *J* = 19.5 Hz).

¹⁹**F NMR** (376 MHz, CDCl₃) δ -81.0 (t, J = 9.6 Hz, 3F), -108.5 (t, J = 12.5 Hz, 2F), -124.2 (s, 2F), -125.6 ~ -125.7 (m, 2F).

HRMS (ESI) m/z calcd. for $C_{26}H_{15}ONBrF_9[M-H]^{-} 643.9867$, found 643.9871.



(*S*)-4-(1-(5-bromo-1*H*-indol-3-yl)-3,3,4,4,5,5,6,6,6-nonafluor o-1-(4-(trifluoromethyl)phenyl)hexyl)phenol (4E)

The reaction was conducted on a 0.1 mmol scale according to the above general procedure A. The product was purified by silica gel flash column chromatography (n-hexane/ethyl acetate

= 100/0 to 2/1) to afford desired product **4E** (54.8 mg, 56 %) as sticky light yellow oil. **HPLC** analysis: Chiralcel AD3 (*n*-hexane/*i*-PrOH = 85/15, flow rate 0.3 mL/min, λ = 234 nm), *t*_R (minor) = 14.54 min, *t*_R (major) = 18.63 min.

¹**H** NMR (500 MHz, CDCl₃) δ 8.26 (s, 1H), 7.67-7.48 (m, 4H), 7.33-7.17 (m, 4H), 7.07 (s, 2H), 6.79 (d, J = 8.7 Hz, 2H), 5.14 (s, 1H), 3.68-3.38 (m, 2H).

¹³**C** NMR (126 MHz, CDCl₃) δ 154.3, 148.8, 136.0, 135.4, 129.5, 128.8 (q, *J* = 32.6 Hz), 128.4, 127.8, 125.5, 125.2, 125.1 (q, *J* = 3.6 Hz), 124.1 (q, *J* = 272.7 Hz), 123.9, 119.6, 115.1, 112.92 (d, *J* = 1.8 Hz), 120.8-106.4 (m), 49.2, 39.0 (t, *J* = 19.5 Hz).

¹⁹**F NMR** (376 MHz, CDCl₃) δ -62.5 (s, 3F), -81.0 (t, J = 9.7 Hz, 3F), -107.9 ~ -109.3 (m, 2F), -124.2 (d, J = 5.4 Hz, 2F), -125.7 (t, J = 12.2 Hz, 2F).

HRMS (ESI) m/z calcd. for C₂₇H₁₅ONBrF₁₂ [M-H]⁻ 676.0151, found 676.0156.



(S)-4-(1-(5-bromo-1*H*-indol-3-yl)-3,3,4,4,5,5,6,6,6-nonafluoro -1-(4-nitrophenyl)hexyl)phenol (4F)

The reaction was conducted on a 0.1 mmol scale according to the above general procedure A. The product was purified by silica gel flash column chromatography (n-hexane/ethyl acetate = 100/0 to 2/1) to afford desired product 4F (62.8 mg, 95 %) as a yellow solid.

HPLC analysis: Chiralcel AD3 (*n*-hexane/*i*-PrOH = 85/15, flow rate 0.5 mL/min, λ = 230 nm), $t_{\rm R}$ (minor) = 15.21 min, $t_{\rm R}$ (major) = 22.34 min.

¹**H NMR** (400 MHz, CDCl₃) δ 8.28 (s, 1H), 8.16 (d, J = 8.9 Hz, 2H), 7.61 (d, J = 8.8Hz, 2H), 7.30-7.22 (m, 4H), 7.09 (s, 1H), 7.03 (s, 1H), 6.79 (d, J = 8.7 Hz, 2H), 4.99 (br s, 1H), 3.55 (t, J = 18.3 Hz, 2H).

¹³C NMR (126 MHz, CDCl₃) δ 154.6, 152.1, 146.3, 135.5, 135.3, 129.5, 129.1, 127.6, 125.4, 125.4, 123.7, 123.4, 119.1, 115.3, 113.1, 120.7-106.0 (m), 49.4, 39.1 (t, J =19.5 Hz).

¹⁹**F NMR** (376 MHz, CDCl₃) δ -81.0 (t, J = 9.1 Hz, 3F), -108.4 (s, 2F), -124.2 (s, 2F), -125.6 (t, J = 12.9 Hz, 2F).

HRMS (ESI) m/z calcd. for C₂₆H₁₅O₃N₂BrF₉ [M-H]⁻ 653.0128, found 653.0139.



(S)-methyl 4-(1-(5-bromo-1H-indol-3-yl)-3,3,4,4,5,5,6,6,6 -nonafluoro-1-(4-hydroxyphenyl)hexyl)benzoate (4G)

The reaction was conducted on a 0.1 mmol scale according to the above general procedure A. The product was purified by silica gel flash column chromatography (n-hexane/ethyl

acetate = 100/0 to 2/1) to afford desired product 4G (46.0 mg, 69 %) as sticky light yellow oil.

HPLC analysis: Chiralcel IA (*n*-hexane/*i*-PrOH = 85/15, flow rate 0.3 mL/min, λ = 230 nm), $t_{\rm R}$ (minor) = 23.38 min, $t_{\rm R}$ (major) = 29.49 min.

¹**H NMR** (500 MHz, CDCl₃) δ 8.24 (s, 1H), 7.94 (d, J = 8.6 Hz, 2H), 7.47 (d, J = 8.5Hz, 2H), 7.25-7.20 (m, 4H), 7.08-6.99 (m, 2H), 6.75 (d, *J* = 8.8 Hz, 2H), 5.17 (s, 1H), 3.89 (s, 3H), 3.60-3.40 (m, 2H).

¹³C NMR (126 MHz, CDCl₃) δ 166.9, 154.3, 150.0, 135.9, 135.4, 129.6, 129.4, 128.3, 128.1, 127.9, 125.5, 125.1, 123.9, 119.6, 115.1, 112.9, 120.6-106.4 (m), 52.2, 49.3, 38.95 (t, J = 19.3 Hz).

¹⁹F NMR (376 MHz, CDCl₃) δ -81.03 (t, J = 9.5 Hz, 3F), -107.80 ~ -109.33 (m, 2F), -124.20 (s, 2F), -125.13 ~ -126.30 (m, 2F).

HRMS (ESI) m/z calcd. for C₂₈H₁₈O₃NBrF₉ [M-H]⁻ 666.0332, found 666.0345.



(S)-4-(1-([1,1'-biphenyl]-4-yl)-1-(5-bromo-1*H*-indol-3-yl)-3,3 ,4,4,5,5,6,6,6-nonafluorohexyl)phenol (4H)

The reaction was conducted on a 0.1 mmol scale according to the above general procedure A. The product was purified by silica gel flash column chromatography (*n*-hexane/ethyl acetate

= 100/0 to 2/1) to afford desired product 4H (39.7 mg, 58 %) as sticky light yellow oil.

HPLC analysis: Chiralcel OD3 (hexane/*i*-PrOH = 87/13, flow rate 0.3 mL/min, $\lambda = 230$ nm), t_R (major) = 25.23 min, t_R (minor) = 28.12 min.

¹**H** NMR (500 MHz, CDCl₃) δ 8.16 (s, 1H), 7.58 (d, *J* = 7.7 Hz, 2H), 7.51 (d, *J* = 8.4 Hz, 2H), 7.42 (dd, *J* = 15.2, 7.9 Hz, 4H), 7.32 (t, *J* = 7.4 Hz, 1H), 7.24 (dt, *J* = 11.6, 8.6 Hz, 4H), 7.11 (s, 1H), 7.08 (d, *J* = 1.3 Hz, 1H), 6.76 (d, *J* = 8.7 Hz, 2H), 4.92 (br s, 1H), 3.63-3.42 (m, 2H).

¹³**C NMR** (126 MHz, CDCl₃) δ 154.1, 144.0, 140.4, 139.1, 136.7, 135.4, 129.7, 128.8, 128.4, 128.1, 127.3, 127.0, 126.7, 125.5, 124.9, 124.2, 120.3, 114.9, 112.8, 120.8-106.3 (m), 48.9, 39.1 (t, *J* = 19.4 Hz).

¹⁹**F NMR** (376 MHz, CDCl₃) δ -80.99 (t, J = 9.6 Hz, 3F), -108.58 (s, 2F), -123.76 ~ -124.57 (m, 2F), -125.31 ~ -126.13 (m, 2F).

HRMS (ESI) m/z calcd. for C₃₂H₂₀ONBrF₉[M-H]⁻ 684.0590, found 684.0602.



(*S*)-4-(1-(5-bromo-1*H*-indol-3-yl)-1-(4-ethynylphenyl)-3,3,4, 4,5,5,6,6,6-nonafluorohexyl)phenol (4I)

The reaction was conducted on a 0.1 mmol scale according to the above general procedure A. The product was purified by silica gel flash column chromatography (*n*-hexane/ethyl acetate = 100/0 to 2/1) to afford desired product **4I** (39.9 mg, 63 %) as

sticky light yellow oil.

HPLC analysis: Chiralcel IA (*n*-hexane/*i*-PrOH = 85/15, flow rate 0.3 mL/min, λ = 230 nm), t_R (minor) = 19.77 min, t_R (major) = 22.39 min.

¹**H** NMR (500 MHz, CDCl₃) δ 8.21 (s, 1H), 7.43 (d, J = 8.4 Hz, 2H), 7.37 (d, J = 8.4 Hz, 2H), 7.23 (d, J = 8.6 Hz, 4H), 7.08 (s, 1H), 7.05 (d, J = 2.0 Hz, 1H), 6.76 (d, J = 8.7 Hz, 2H), 3.50 (t, J = 18.7 Hz, 2H), 3.08 (s, 1H).

¹³**C NMR** (126 MHz, CDCl₃) δ 154.2, 145.6, 136.2, 135.4, 131.9, 129.6, 128.1, 128.0, 125.4, 125.0, 124.0, 120.3, 119.8, 115.0, 112.9, 112.8, 120.8-106.5 (m), 83.4, 77.5, 49.1, 39.0 (t, *J* = 19.4 Hz).

¹⁹**F NMR** (376 MHz, CDCl₃) δ -81.0 (t, J = 9.6 Hz, 3F), -108.6 (d, J = 8.3 Hz, 2F), -124.2 (s, 2F), -125.7 (t, J = 13.4 Hz, 2F).

HRMS (ESI) m/z calcd. for C₂₈H₁₆ONBrF₉[M-H]⁻ 632.0277, found 632.0288.



(*S*)-4-(1-(5-bromo-1*H*-indol-3-yl)-3,3,4,4,5,5,6,6,6-nonafluor o-1-(3-methoxyphenyl)hexyl)phenol (4J)

The reaction was conducted on a 0.1 mmol scale according to the above general procedure A. The product was purified by silica gel flash column chromatography (n-hexane/ethyl acetate

= 100/0 to 2/1) to afford desired product 4J (47.9 mg, 75 %) as sticky light yellow oil. HPLC analysis: Chiralcel AD3 (*n*-hexane/*i*-PrOH = 85/15, flow rate 0.3 mL/min, λ = 230 nm), $t_{\rm R}$ (minor) = 18.32 min, $t_{\rm R}$ (major) = 20.30 min.

¹**H NMR** (500 MHz, CDCl₃) δ 8.16 (s, 1H), 7.34-7.14 (m, 5H), 7.11 (s, 1H), 7.06 (s, 1H), 7.03-6.90 (m, 2H), 6.76 (t, *J* = 8.9 Hz, 3H), 3.74 (s, 3H), 3.59-3.42 (m, 2H).

¹³**C NMR** (126 MHz, CDCl₃) δ 159.2, 154.1, 146.9, 136.5, 135.4, 129.6, 129.0, 128.2, 125.6, 124.9, 124.21, 120.5, 120.11, 114.8, 114.75, 112.7, 112.7, 111.2, 119.9-106.4 (m), 55.2, 49.1, 39.01 (t, *J* = 19.4 Hz).

¹⁹**F NMR** (376 MHz, CDCl₃) δ -81.03 (s, 3F), -108.63 (s, 2F), -124.21 (s, 2F), -125.63 (s, 2F).

HRMS (ESI) m/z calcd. for C₂₇H₁₈O₂NBrF₉ [M-H]⁻ 638.03827, found 638.03784.



(*S*)-4-(1-(5-bromo-1*H*-indol-3-yl)-3,3,4,4,5,5,6,6,6-nonafluoro -1-(2-fluorophenyl)hexyl)phenol (4K)

The reaction was conducted on a 0.1 mmol scale according to the above general procedure A. The product was purified by silica gel flash column chromatography (n-hexane/ethyl acetate =

100/0 to 2/1) to afford desired product **4K** (50.1 mg, 80 %) as sticky light yellow oil. **HPLC** analysis: Chiralcel IA (*n*-hexane/*i*-PrOH = 85/15, flow rate 0.3 mL/min, λ = 230 nm), t_R (minor) = 17.89 min, t_R (major) = 21.70 min.

¹**H NMR** (500 MHz, CDCl₃) δ 8.18 (s, 1H), 7.38-7.30 (m, 1H), 7.27-7.15 (m, 6H), 7.13-6.99 (m, 3H), 6.80-6.69 (m, 2H), 4.78 (br s, 1H), 3.72-3.53 (m, 2H).

¹³**C** NMR (126 MHz, CDCl₃) δ 161.6 (d, J = 248.3 Hz), 154.0, 135.9, 135.3, 131.4 (d, J = 3.8 Hz), 129.6 (d, J = 8.9 Hz), 129.3 (d, J = 11.2 Hz), 129.1, 128.1, 125.0, 124.2 (d, J = 5.3 Hz), 124.0, 123.8 (d, J = 3.0 Hz), 120.0, 116.4, 116.2, 114.8, 112.73, 112.70, 120.8-106.5 (m), 48.0, 38.8 (td, J = 19.4, 4.7 Hz).

¹⁹**F** NMR (376 MHz, CDCl₃) δ -81.0 (t, J = 9.5 Hz, 3F), -104.0 (s, 1F), -107.5 ~ -110.7 (m, 2F), -124.3 (d, J = 7.5 Hz, 2F), -125.6 (dt, J = 27.7, 14.6 Hz, 2F). HRMS (ESI) m/z calcd. for C₂₆H₁₅ONBrF₁₀ [M-H]⁻ 626.0183, found 626.0192.



(*S*)-4-(1-(5-bromo-1*H*-indol-3-yl)-3,3,4,4,5,5,6,6,6-nonafluoro -1-(thiophen-2-yl)hexyl)phenol (4L)

The reaction was conducted on a 0.1 mmol scale according to the above general procedure A. The product was purified by silica gel flash column chromatography (n-hexane/ethyl acetate =

100/0 to 2/1) to afford desired product 4L (58.3 mg, 95 %) as sticky light yellow oil. HPLC analysis: Chiralcel AD3 (*n*-hexane/*i*-PrOH = 85/15, flow rate 0.3 mL/min, λ = 230 nm), *t*_R (minor) = 18.63 min, *t*_R (major) = 24.27 min.

¹**H** NMR (500 MHz, CDCl₃) δ 8.14 (s, 1H), 7.25-7.18 (m, 5H), 7.07 (d, J = 8.2 Hz, 2H), 7.02 (s, 1H), 6.95 (d, J = 5.0 Hz, 1H), 6.74 (d, J = 8.6 Hz, 2H), 5.00 (br s, 1H), 3.60-3.25 (m, 2H).

¹³**C NMR** (126 MHz, CDCl₃) δ 154.2, 146.4, 136.4, 135.4, 129.3, 128.6, 127.9, 125.2, 125.0, 124.9, 124.0, 121.7, 120.5, 114.9, 112.7, 120.6-106.6 (m), 46.8, 39.8 (t, *J* = 19.8 Hz).

¹⁹**F** NMR (376 MHz, CDCl₃) δ -81.0 (t, J = 9.5 Hz, 3F), -106.9 ~ -110.6 (m, 2F), -124.3 (d, J = 7.6 Hz, 2F), -125.7 (s, 2F).

HRMS (ESI) m/z calcd. for C₂₄H₁₄ONBrF₉S [M-H]⁻ 613.9841, found 613.9846.



(*S*)-4-(1-(5-bromo-1*H*-indol-3-yl)-3,3,4,4,5,5,6,6,6-nonafluoro -1-(naphthalen-2-yl)hexyl)phenol (4M)

The reaction was conducted on a 0.1 mmol scale according to the above general procedure A. The product was purified by silica gel flash column chromatography (n-hexane/ethyl acetate =

100/0 to 2/1) to afford desired product **4M** (62.0 mg, 94 %) as sticky light yellow oil. **HPLC** analysis: Chiralcel AD3 (*n*-hexane/*i*-PrOH = 85/15, flow rate 0.4 mL/min, λ = 230 nm), *t*_R (minor) = 15.90 min, *t*_R (major) = 24.35 min.

¹**H NMR** (500 MHz, CDCl₃) δ 8.18 (s, 1H), 7.90 (s, 1H), 7.80 (s, 2H), 7.74 (d, *J* = 8.7 Hz, 1H), 7.49 (d, *J* = 6.6 Hz, 3H), 7.30 (dd, *J* = 9.5, 5.0 Hz, 2H), 7.25 (d, *J* = 8.8 Hz, 2H), 7.18 (s, 1H), 7.06 (s, 1H), 6.76 (d, *J* = 8.4 Hz, 2H), 3.76-3.55 (m, 2H).

¹³**C NMR** (126 MHz, CDCl₃) δ 154.1, 142.3, 136.7, 135.4, 133.0, 132.0, 129.8, 128.4, 128.2, 127.6, 127.4, 127.1, 126.2, 125.9, 125.8, 124.9, 124.2, 120.0, 115.0, 112.8, 112.8, 120.5-106.5 (m), 49.3, 39.0 (t, *J* = 19.5 Hz).

¹⁹**F NMR** (376 MHz, CDCl₃) δ -81.0 (t, J = 9.6 Hz, 3F), -107.4 ~ -109.7 (m, 2F), -124.1 (d, J = 9.2 Hz, 2F), -125.6 (dd, J = 23.1, 11.0 Hz, 2F).

HRMS (ESI) m/z calcd. for $C_{30}H_{18}ONBrF_9 [M-H]^- 658.0434$, found 658.0441.



(*S*)-4-(1-(5-bromo-1*H*-indol-3-yl)-3,3,4,4,5,5,6,6,6-nonafluor o-1-(*p*-tolyl)hexyl)-2-fluorophenol (4N)

The reaction was conducted on a 0.1 mmol scale according to the above general procedure A. The product was purified by silica gel flash column chromatography (n-hexane/ethyl acetate

= 100/0 to 2/1) to afford desired product **4N** (62.2 mg, 97 %) as sticky light yellow oil. **HPLC** analysis: Chiralcel AD3 (*n*-hexane/*i*-PrOH = 85/15, flow rate 0.3 mL/min, λ = 240 nm), *t*_R (minor) = 15.46 min, *t*_R (major) = 18.98 min.

¹**H NMR** (500 MHz, CDCl₃) δ 8.17 (s, 1H), 7.33-7.19 (m, 4H), 7.16-7.01 (m, 6H), 6.90 (t, *J* = 8.8 Hz, 1H), 5.27 (s, 1H), 3.47 (t, *J* = 18.2 Hz, 2H), 2.31 (s, 3H).

¹³**C** NMR (126 MHz, CDCl₃) δ 150.4 (d, J = 236.9 Hz), 142.0 (d, J = 14.3 Hz), 141.4, 137.8 (d, J = 4.8 Hz), 136.4, 135.4, 128.9, 128.0, 127.8, 125.5, 125.0, 124.5 (d, J = 2.6 Hz), 124.0, 119.9, 116.6 (d, J = 1.8 Hz), 115.9 (d, J = 19.8 Hz), 112.81, 112.77, 120.8-106.5 (m), 48.8, 39.1 (t, J = 19.4 Hz), 20.9.

¹⁹**F NMR** (376 MHz, CDCl₃) δ -81.0 (t, *J* = 8.5 Hz, 3F), -108.6 (d, *J* = 14.6 Hz, 2F), -124.2 (s, 2F), -125.6 (t, *J* = 13.0 Hz, 2F), -138.6 ~ -141.1 (m, 1F).

HRMS (ESI) m/z calcd. for $C_{27}H_{17}ONBrF_{10}[M-H]^{-}$ 640.0339, found 640.0350.



(*S*)-4-(1-(5-bromo-1*H*-indol-3-yl)-3,3,4,4,5,5,6,6,6-nonafluoro -1-(*p*-tolyl)hexyl)-2-methylphenol (4O)

The reaction was conducted on a 0.1 mmol scale according to the above general procedure A. The product was purified by silica gel flash column chromatography (n-hexane/ethyl acetate =

100/0 to 2/1) to afford desired product **4O** (44.6 mg, 70 %) as sticky light yellow oil. **HPLC** analysis: Chiralcel AD3 (*n*-hexane/*i*-PrOH = 85/15, flow rate 0.3 mL/min, λ = 230 nm), *t*_R (minor) = 15.72 min, *t*_R (major) = 18.65 min. ¹**H NMR** (500 MHz, CDCl₃) δ 8.11 (s, 1H), 7.27-7.22 (m, 2H), 7.20 (s, 2H), 7.12-6.99 (m, 6H), 6.63 (d, J = 8.4 Hz, 1H), 4.80 (s, 1H), 3.58-3.38 (m, 2H), 2.30 (s, 3H), 2.16 (s, 3H).

¹³**C NMR** (126 MHz, CDCl₃) δ 152.2, 142.1, 137.0, 136.0, 135.4, 130.8, 128.7, 128.3, 127.9, 126.9, 125.6, 124.8, 124.3, 123.1, 120.5, 114.3, 112.7, 112.6, 120.7-106.6 (m), 48.7, 39.0 (t, *J* = 19.6 Hz), 20.9, 16.1.

¹⁹**F NMR** (376 MHz, CDCl₃) δ -81.0 (t, J = 9.7 Hz, 3F), -108.7 (t, J = 13.0 Hz, 2F), -124.2 (d, J = 4.8 Hz, 2F), -125.2 ~ -126.4 (m, 2F).

HRMS (ESI) m/z calcd. for C₂₈H₂₀ON BrF₉ [M-H]⁻ 636.0590, found 636.0605.



(*S*)-4-(1-(5-bromo-1*H*-indol-3-yl)-3,3,4,4,5,5,6,6,6-nonafluor o-1-(*p*-tolyl)hexyl)-3-fluorophenol (4P)

The reaction was conducted on a 0.1 mmol scale according to the above general procedure A. The product was purified by silica gel flash column chromatography (n-hexane/ethyl acetate

= 100/0 to 2/1) to afford desired product **4P** (57.1 mg, 89 %) as sticky light yellow oil. **HPLC** analysis: Chiralcel AD3 (*n*-hexane/*i*-PrOH = 85/15, flow rate 0.4 mL/min, λ = 240 nm), $t_{\rm R}$ (minor) = 11.83 min, $t_{\rm R}$ (major) = 22.86 min.

¹**H** NMR (500 MHz, CDCl₃) δ 8.14 (s, 1H), 7.22-7.12 (m, 5H), 7.06-7.05 (m, 3H), 7.00 (t, *J* = 8.9 Hz, 1H), 6.60 (dd, *J* = 13.1, 2.5 Hz, 1H), 6.46 (dd, *J* = 8.7, 2.5 Hz, 1H), 3.73-3.57 (m, 1H), 3.57-3.38 (m, 1H), 2.28 (s, 3H).

¹³**C** NMR (126 MHz, CDCl₃) δ 162.0 (d, J = 248.6 Hz), 156.5 (d, J = 12.1 Hz), 141.1, 136.2, 135.3, 132.3 (d, J = 5.9 Hz), 128.7, 128.1, 127.4, 124.9, 124.1, 123.9 (d, J = 5.9 Hz), 121.1 (d, J = 12.0 Hz), 120.3, 112.7, 110.6 (d, J = 2.4 Hz), 104.1, 103.9, 120.7-106.5 (m), 47.6, 38.6 (t, J = 19.4 Hz). 20.9.

¹⁹**F NMR** (376 MHz, CDCl₃) δ -81.0 (t, J = 9.2 Hz, 3F), -101.8 (s, 1F), -109.4 (dd, J = 1000.5, 275.3 Hz, 2F), -124.3 (d, J = 6.4 Hz, 2F), -125.3 ~ -126.1 (m, 2F). **HRMS** (ESI) m/z calcd. for C₂₇H₁₇ON BrF₁₀ [M-H]⁻ 640.0339, found 640.0347.



(*S*)-4-(3,3,4,4,5,5,6,6,6-nonafluoro-1-(1*H*-indol-3-yl)-1-(*p*-toly l)hexyl)phenol (4Q)

The reaction was conducted on a 0.1 mmol scale according to the above general procedure A. The product was purified by silica gel flash column chromatography (n-hexane/ethyl acetate =

100/0 to 2/1) to afford desired product **4Q** (51.8 mg, 95 %) as sticky light yellow oil. **HPLC** analysis: Chiralcel AD3 (*n*-hexane/*i*-PrOH = 90/10, flow rate 0.2 mL/min, λ = 230 nm), t_R (major) = 40.11 min, t_R (minor) = 42.34 min.

¹**H** NMR (500 MHz, CDCl₃) δ 8.02 (s, 1H), 7.36-7.31 (m, 3H), 7.29 (d, J = 8.7 Hz, 2H), 7.17-7.13 (m, 1H), 7.09 (d, J = 8.2 Hz, 2H), 7.03 (d, J = 8.1 Hz, 1H), 7.00 (d, J = 2.3 Hz, 1H), 6.97-6.90 (m, 1H), 6.74-6.65 (m, 2H), 3.56 (t, J = 18.6 Hz, 2H), 2.32 (s, 3H).

¹³**C NMR** (126 MHz, CDCl₃) δ 153.8, 142.4, 137.3, 136.8, 135.9, 129.8, 128.7, 128.1, 126.4, 124.1, 122.0, 121.8, 121.1, 119.2, 114.7, 111.3, 120.9-106.5 (m), 49.0, 39.2 (t, *J* = 19.4 Hz), 20.9.

¹⁹**F NMR** (376 MHz, CDCl₃) δ -81.0 (t, J = 9.8 Hz, 3F), -108.5 (t, J = 12.9 Hz, 2F), -123.6 ~ -124.7 (m, 2F), -125.0 ~ -126.1 (m, 2F). **HRMS** (ESI) m/z calcd. for C₂₇H₁₉ONF₉ [M-H]⁻ 544.1328, found 544.1334.



(*S*)-4-(3,3,4,4,5,5,6,6,6-nonafluoro-1-(5-iodo-1*H*-indol-3-yl)-1-(*p*-tolyl)hexyl)phenol (4R)

The reaction was conducted on a 0.1 mmol scale according to the above general procedure A. The product was purified by silica gel flash column chromatography (*n*-hexane/ethyl acetate = 10^{-10} flash column chromatography (*n*-hexane

100/0 to 2/1) to afford desired product **4R** (63.8 mg, 95 %) as sticky light yellow oil. **HPLC** analysis: Chiralcel IA (*n*-hexane/*i*-PrOH = 85/15, flow rate 0.25 mL/min, λ = 230 nm), *t*_R (minor) = 21.62 min, *t*_R (major) = 24.75 min.

¹**H** NMR (500 MHz, CDCl₃) δ 8.12 (s, 1H), 7.37 (dd, J = 8.5, 1.4 Hz, 1H), 7.30-7.17 (m, 5H), 7.12 (d, J = 8.5 Hz, 1H), 7.07 (d, J = 8.2 Hz, 2H), 6.99 (d, J = 2.1 Hz, 1H), 6.73 (d, J = 8.7 Hz, 2H), 3.58-3.35 (m, 2H), 2.31 (s, 3H).

¹³C NMR (126 MHz, CDCl₃) δ 153.9, 142.0, 136.9, 136.1, 135.8, 130.5, 130.3, 129.6, 129.0, 128.7, 127.9, 124.9, 120.3, 114.8, 113.2, 120.5-106.2 (m), 83.0, 48.8, 39.2 (t, *J* = 19.4 Hz), 20.9.

¹⁹**F NMR** (376 MHz, CDCl₃) δ -81.0 (t, J = 9.7 Hz, 3F), -108.6 (d, J = 10.5 Hz, 2F), -124.2 (dd, J = 12.0, 7.3 Hz, 2F), -125.2 ~ -126.1 (m, 2F).

HRMS (ESI) m/z calcd. for C₂₇H₁₈ONIF₉ [M-H]⁻ 670.0295, found 670.0314.



(*S*)-4-(1-(5-chloro-1*H*-indol-3-yl)-3,3,4,4,5,5,6,6,6-nonafluor o-1-(*p*-tolyl)hexyl)phenol (4S)

The reaction was conducted on a 0.1 mmol scale according to the above general procedure A. The product was purified by silica gel flash column chromatography (n-hexane/ethyl acetate

= 100/0 to 2/1) to afford desired product **4S** (53.3 mg, 92 %) as sticky light yellow oil. **HPLC** analysis: Chiralcel IA (*n*-hexane/*i*-PrOH = 85/15, flow rate 0.25 mL/min, λ = 240 nm), *t*_R (minor) = 20.94 min, *t*_R (major) = 23.26 min.

¹**H NMR** (500 MHz, CDCl₃) δ 8.15 (s, 1H), 7.29-7.25 (m, 5H), 7.16-7.04 (m, 4H), 6.94 (s, 1H), 6.75 (d, *J* = 8.6 Hz, 2H), 5.00 (s, 1H), 3.64-3.36 (m, 2H), 2.33 (s, 3H).

¹³C NMR (126 MHz, CDCl₃) δ 154.0, 142.0, 136.9, 136.1, 135.1, 129.6, 128.8, 127.9, 127.5, 125.5, 124.9, 122.3, 121.2, 120.6, 114.8, 112.2, 120.6-106.6 (m), 48.8, 39.1 (t, *J* = 19.5 Hz), 20.9.

¹⁹**F NMR** (376 MHz, CDCl₃) δ -81.0 (t, J = 9.8 Hz, 3F), -108.6 (d, J = 14.6 Hz, 2F), -124.2 (d, J = 3.2 Hz, 2F), -125.3 ~ -126.2 (m, 2F).

HRMS (ESI) m/z calcd. for C₂₇H₁₈ONClF₉[M-H]⁻ 578.0939, found 578.0948.



(S)-4-(3,3,4,4,5,5,6,6,6-nonafluoro-1-(5-fluoro-1*H*-indol-3-yl)-1-(*p*-tolyl)hexyl)phenol (4T)

The reaction was conducted on a 0.1 mmol scale according to the above general procedure A. The product was purified by silica gel flash column chromatography (n-hexane/ethyl acetate =

100/0 to 2/1) to afford desired product **4T** (36.1 mg, 56 %) as sticky light yellow oil. **HPLC** analysis: Chiralcel IA (*n*-hexane/*i*-PrOH = 85/15, flow rate 0.2 mL/min, λ = 230 nm), $t_{\rm R}$ (minor) = 26.49 min, $t_{\rm R}$ (major) = 28.80 min.

¹**H NMR** (500 MHz, CDCl₃) δ 8.09 (s, 1H), 7.33-7.18 (m, 6H), 7.12-7.02 (m, 3H), 6.87 (td, J = 8.9, 1.9 Hz, 1H), 6.72 (d, J = 8.6 Hz, 2H), 6.60 (dd, J = 10.2, 1.2 Hz, 1H), 4.95 (br s, 1H), 3.48 (t, J = 18.6 Hz, 2H), 2.30 (s, 3H).

¹³C NMR (126 MHz, CDCl₃) δ 157.1 (d, J = 234.0 Hz), 153.9, 142.0, 136.9, 136.0, 133.3, 129.6, 128.7, 127.9, 126.8 (d, J = 10.0 Hz), 125.8, 121.1 (d, J = 4.6 Hz), 114.8, 111.8 (d, J = 9.8 Hz), 110.3 (d, J = 26.5 Hz), 106.8 (d, J = 24.3 Hz), 120.8-106.4 (m), 48.8, 39.1 (t, J = 19.5 Hz), 20.9.

¹⁹**F NMR** (376 MHz, CDCl₃) δ -80.4 ~ -81.4 (m, 3F), -108.4 ~ -108.9 (m, 2F), -123.9 ~ -124.1 (m, 1F), -124.3 (s, 2F), -125.5 ~ -126.0 (m, 2F).

HRMS (ESI) m/z calcd. for C₂₇H₁₈ONBrF₁₀ [M-H]⁻ 562.1234, found 562.1248.



(*S*)-4-(3,3,4,4,5,5,6,6,6-nonafluoro-1-(5-methyl-1*H*-indol-3-y l)-1-(*p*-tolyl)hexyl)phenol (4U)

The reaction was conducted on a 0.1 mmol scale according to the above general procedure A. The product was purified by silica gel flash column chromatography (n-hexane/ethyl acetate

= 100/0 to 2/1) to afford desired product 4U (53.7 mg, 96 %) as sticky light yellow oil. HPLC analysis: Chiralcel AD3 (*n*-hexane/*i*-PrOH = 85/15, flow rate 0.3 mL/min, λ = 214 nm), t_R (minor) = 17.74 min, t_R (major) = 21.31 min.

¹**H NMR** (500 MHz, CDCl₃) δ 7.97 (s, 1H), 7.30-7.23 (m, 5H), 7.07 (d, *J* = 7.9 Hz, 2H), 6.97-6.94 (m, 2H), 6.77 (s, 1H), 6.71 (d, *J* = 8.3 Hz, 2H), 4.95 (s, 1H), 3.63-3.40 (m, 2H), 2.31 (s, 3H), 2.28 (s, 3H).

¹³C NMR (126 MHz, CDCl₃) δ 153.8, 142.4, 137.4, 135.8, 135.2, 129.9, 128.6, 128.3, 128.1, 126.6, 124.3, 123.5, 121.6, 120.5, 114.6, 110.9, 120.7-106.5 (m), 49.1, 39.1 (t, *J* = 19.4 Hz), 21.7, 20.9.

¹⁹**F NMR** (376 MHz, CDCl₃) δ -81.0 (t, J = 9.4 Hz, 3F), -108.3 (s, 2F), -124.2 (s, 2F), -125.0 ~ -126.1 (m, 2F).

HRMS (ESI) m/z calcd. for C₂₈H₂₁ONF₉[M-H]⁻ 558.1485, found 558.1490.



(*S*)-4-(3,3,4,4,5,5,6,6,6-nonafluoro-1-(5-methoxy-1*H*-indol-3 -yl)-1-(*p*-tolyl)hexyl)phenol (4V)

The reaction was conducted on a 0.1 mmol scale according to the above general procedure A. The product was purified by silica gel flash column chromatography (n-hexane/ethyl acetate

= 100/0 to 2/1) to afford desired product 4V (54.6 mg, 56 %) as sticky light yellow oil. HPLC analysis: Chiralcel AD3 (*n*-hexane/*i*-PrOH = 85/15, flow rate 0.3 mL/min, λ = 214 nm), t_R (minor) = 18.25 min, t_R (major) = 23.57 min.

¹**H** NMR (500 MHz, CDCl₃) δ 7.98 (s, 1H), 7.34-7.24 (m, 4H), 7.22 (d, J = 8.8 Hz, 1H), 7.06 (d, J = 8.2 Hz, 2H), 7.01 (d, J = 2.3 Hz, 1H), 6.79 (dd, J = 8.8, 2.4 Hz, 1H), 6.70 (d, J = 8.7 Hz, 2H), 6.37 (d, J = 2.0 Hz, 1H), 5.05 (s, 1H), 3.58 (s, 3H), 3.54-3.42 (m, 2H), 2.29 (s, 3H).

¹³**C NMR** (126 MHz, CDCl₃) δ 153.9, 153.2, 142.2, 137.0, 135.9, 132.0, 129.8, 128.6, 128.1, 126.9, 124.8, 120.7, 114.6, 111.8, 111.7, 104.4, 120.6-106.6 (m), 55.7, 48.9, 39.3 (t, *J* = 19.4 Hz), 20.9.

¹⁹**F NMR** (376 MHz, CDCl₃) δ -81.0 (t, J = 9.5 Hz, 3F), -108.3 (s, 2F), -124.3 (s, 2F), -125.1 ~ -126.2 (m, 2F).

HRMS (ESI) m/z calcd. for C₂₈H₂₁O₂NF₉ [M-H]⁻ 574.1434, found 574.1441.



(*R*)-methyl 4-(5-bromo-1*H*-indol-3-yl)-2,2-difluoro-4-(4 -hydroxy-phenyl)-4-(*p*-tolyl)butanoate (5A)

The reaction was conducted on a 0.025 mmol scale according to the above general procedure A. The product was purified by silica gel flash column chromatography (*n*-hexane/ethyl

acetate = 100/0 to 2/1) to afford desired product **5A** (5.4 mg, 42 %) as a sticky light yellow solid.

HPLC analysis: Chiralcel IC (*n*-hexane/*i*-PrOH = 85/15, flow rate 0.2 mL/min, λ = 240 nm), t_R (major) = 35.06 min, t_R (minor) = 38.24 min.

¹**H** NMR (500 MHz, CDCl₃) δ 8.34 (s, 1H), 7.31-7.28 (m, 4H), 7.21 (s, 2H), 7.10-7.03 (m, 3H), 7.01 (d, J = 2.4 Hz, 1H), 6.72 (d, J = 8.7 Hz, 2H), 4.88 (s, 1H), 3.76 (t, J = 14.7 Hz, 2H), 2.99 (s, 3H), 2.31 (s, 3H).

¹³**C** NMR (126 MHz, CDCl₃) δ 163.1 (t, J = 33.4 Hz), 153.7, 142.8, 137.9, 135.8, 135.2, 129.2, 128.8, 128.0, 127.7, 127.5, 124.6, 124.4, 119.2, 115.6 (d, J = 251.3 Hz), 114.8, 112.8, 112.5, 52.5, 48.2, 42.9 (t, J = 22.7 Hz), 20.9.

¹⁹**F NMR** (376 MHz, CDCl₃) δ -97.6 ~ -99.4 (m, 2F).

HRMS (ESI) m/z calcd. for C₂₆H₂₁O₃NBrF₂ [M-H]⁻ 514.06469, found 514.06525.

General procedure B: Cu/CPA-catalyzed asymmetric three-component radical-initiated trichloromethylarylation of 1,1-diarylalkene



Under argon atmosphere, an oven-dried resealable Schlenk tube equipped with a magnetic stir bar was charged with 1,1-diarylalkene substrate **1** (0.1 mmol, 1.0 equiv.), substituted indole (0.12 mmol, 1.2 equiv.), CuI (3.0 mg, 0.015 mmol, 15 mol%), chiral phosphoric acid ((*S*)-A5 (9.86 mg, 0.01 mmol, 10 mol%), CCl₃SO₂Cl (**3c**, 26.2 mg, 0.12 mmol, 1.2 equiv.), Ag₂CO₃ (16.56 mg, 0.06 mmol, 0.6 equiv.), and anhydrous DCM (1.0 mL) at 0 °C, then the sealed tube was stirred at -35 °C for 60-72 h. Upon completion (monitored by TLC), the reaction mixture was directly purified by a silica gel chromatography [eluent: *n*-hexane/ethyl acetate = 100/0 to 2/1, using petroleum ether (100%) to remove the solvent (DCM) at first] to afford the desired product **6**.

Note: the trichloromethyl-containing products **6A–6D** was unstable at 2-30°C, easy to convert to other products, especially when dissolved in halogen-containing solvents, such as DCM, CHCl₃ etc, and slightly more stable in alcoholic solvents, such as *n*-hexane, ethyl acetate, alcohol isopropanol etc. So **6A–6D** should be characterized by NMR, HPLC or HRMS ASAP. Meanwhile, the reaction is sensitive to water and air, Schlenk tube and the reagents must be dried prior to use.



The racemate was prepared following the same procedure described above using CuI (3.0 mg, 0.015 mmol, 15 mol%) and diphenyl phosphate (2.5 mg, 0.01 mmol, 10 mol%) as catalyst at -35 °C in anhydrous DCM (1.0 mL) for 48 h. Upon completion (monitored by TLC), the solvent was removed *in vacuo*, and the residue was purified by a silica gel column chromatography (eluent: *n*-hexane/ethyl acetate = 100/0 to 2/1) to give the desired product.

Note: Since the reaction is sensitive to water and air, Schlenk tube and the reagents must be dried prior to use.



(S)-4-(1-(5-bromo-1*H*-indol-3-yl)-3,3,3-trichloro-1-(*p*-tolyl) propyl)phenol (6A)

The reaction was conducted on a 0.1 mmol scale according to the above general procedure B. The product was purified by silica gel flash column chromatography (*n*-hexane/ethyl acetate = 100/0 to 2/1) to afford desired product **6A** (44.9 mg, 86 %) as sticky light yellow oil.

HPLC analysis: Chiralcel AD3 (*n*-hexane/*i*-PrOH = 85/15, flow rate 0.5 mL/min, λ = 230 nm), t_R (minor) = 16.08 min, t_R (major) = 24.65 min.

¹**H NMR** (400 MHz, MeOD) δ 7.46 (d, J = 8.4 Hz, 2H), 7.42-7.36 (m, 2H), 7.33 (s, 1H), 7.24 (d, J = 8.6 Hz, 1H), 7.12-7.02 (m, 4H), 6.88 (d, J = 1.6 Hz, 1H), 6.72-6.68 (m, 2H), 4.32 (s, 2H), 2.25 (s, 3H).

¹³C NMR (101 MHz, MeOD) δ 155.2, 142.9, 136.1, 135.7, 135.3, 129.4, 128.9, 128.1, 127.7, 127.4, 123.8, 123.4, 117.3, 114.2, 112.6, 111.2, 98.2, 58.8, 51.7, 19.5.

HRMS (ESI) m/z calcd. for $C_{24}H_{18}ON^{81}BrCl_3 [M-H]^- 521.96224$, found 521.96173.



(S)-4-(1-(5-bromo-1*H*-indol-3-yl)-3,3,3-trichloro-1-(2-fluoro phenyl)propyl)phenol (6B)

The reaction was conducted on a 0.1 mmol scale according to the above general procedure B. The product was purified by silica gel flash column chromatography (*n*-hexane/ethyl acetate = 100/0 to 2/1) to afford desired product **6B** (43.1 mg, 82 %) as

sticky light yellow oil.

HPLC analysis: Chiralcel AD3 (*n*-hexane/*i*-PrOH = 85/15, flow rate 0.5 mL/min, λ = 254 nm), t_R (minor) = 20.34 min, t_R (major) = 27.49 min.

¹**H** NMR (400 MHz, MeOD) δ 7.65 (td, J = 8.2, 1.6 Hz, 1H), 7.42 (s, 1H), 7.32 (d, J = 8.0 Hz, 2H), 7.30-7.21 (m, 2H), 7.13-7.04 (m, 2H), 6.99 (dd, J = 13.1, 8.1 Hz, 1H), 6.93 (d, J = 1.6 Hz, 1H), 6.73-6.66 (m, 2H), 4.52 (d, J = 15.8 Hz, 1H), 4.37 (d, J = 15.7 Hz, 1H).

¹³**C NMR** (101 MHz, MeOD) δ 161.5 (d, J = 247.5 Hz), 155.4, 135.7, 135.0, 131.5 (d, J = 4.0 Hz), 130.2 (d, J = 9.9 Hz), 128.9, 128.8, 128.2, 126.5, 123.5 (d, J = 3.4 Hz), 123.1 (d, J = 3.1 Hz), 116.9, 116.2, 116.0, 114.1, 112.7, 111.2, 98.0, 58.9 (d, J = 7.0 Hz), 51.1 (d, J = 2.3 Hz).

¹⁹**F NMR** (376 MHz, MeOD) δ -104.5 (s, 1F).

HRMS (ESI) m/z calcd. for C₂₃H₁₅ON⁸¹BrCl₃F [M-H]⁻ 525.93716, found 525.93707.



(S)-4-(1-(4-(*tert*-butyl)phenyl)-3,3,3-trichloro-1-(5-fluoro-1*H*-indol-3-yl)propyl)phenol (6C)

The reaction was conducted on a 0.1 mmol scale according to the above general procedure B. The product was purified by silica gel flash column chromatography (n-hexane/ethyl acetate =

100/0 to 2/1) to afford desired product **6C** (34.2 mg, 68 %) as sticky light yellow oil. **HPLC** analysis: Chiralcel AD3 (*n*-hexane/*i*-PrOH = 85/15, flow rate 0.5 mL/min, λ = 230 nm), $t_{\rm R}$ (minor) = 15.58 min, $t_{\rm R}$ (major) = 18.58 min.

¹**H NMR** (400 MHz, MeOD) δ 7.48 (s, 1H), 7.45 (s, 1H), 7.35 (s, 1H), 7.33 (s, 1H), 7.30 (s, 1H), 7.23-7.18 (m, 3H), 6.70 (td, J = 9.0, 2.4 Hz, 1H), 6.65-6.63 (m, 1H), 6.62 (s, 1H), 6.29 (dd, J = 10.9, 2.4 Hz, 1H), 4.34-4.18 (m, 2H), 1.21 (s, 9H).

¹³C NMR (101 MHz, MeOD) δ 156.7 (d, J = 231.5 Hz), 155.2, 148.5, 143.1, 136.1, 133.6, 129.5, 127.8, 127.4, 124.4, 117.6 (d, J = 4.7 Hz), 114.1, 111.6 (d, J = 9.8 Hz), 108.7 (d, J = 26.5 Hz), 106.0 (d, J = 24.5 Hz), 98.2, 58.6, 51.6, 33.7, 30.4. ¹⁹F NMR (376 MHz, MeOD) δ -127.2 (s, 1F).

HRMS (ESI) m/z calcd. for C₂₇H₂₄ONCl₃F [M-H]⁻ 502.09130, found 502.09097.



(S)-2-fluoro-4-(3,3,3-trichloro-1-(5-methoxy-1*H*-indol-3-yl)-1-(*p*-tolyl)propyl)phenol (6D)

The reaction was conducted on a 0.1 mmol scale according to the above general procedure B. The product was purified by silica gel flash column chromatography (n-hexane/ethyl acetate

= 100/0 to 2/1) to afford desired product **6D** (41.7 mg, 85 %) as sticky light yellow oil. **HPLC** analysis: Chiralcel AD3 (*n*-hexane/*i*-PrOH = 85/15, flow rate 0.6 mL/min, λ = 230 nm), $t_{\rm R}$ (minor) = 15.81 min, $t_{\rm R}$ (major) = 19.96 min.

¹**H** NMR (500 MHz, CDCl₃) δ 8.12 (s, 1H), 7.47 (d, J = 8.4 Hz, 2H), 7.34 (dd, J = 12.9, 2.3 Hz, 1H), 7.29 (dd, J = 8.0, 2.3 Hz, 1H), 7.24 (t, J = 5.3 Hz, 2H), 7.10 (t, J = 6.8 Hz, 2H), 6.90 (t, J = 8.9 Hz, 1H), 6.81 (dd, J = 8.8, 2.4 Hz, 1H), 6.27 (d, J = 2.3 Hz, 1H), 4.30 (s, 2H), 3.59 (s, 3H), 2.31 (s, 3H).

¹³**C** NMR (126 MHz, CDCl₃) δ 153.2, 150.4 (d, J = 236.5 Hz), 142.2, 141.7 (d, J = 14.4 Hz), 138.3 (d, J = 4.8 Hz), 135.9, 131.8, 128.8, 127.9, 127.5, 126.6, 124.6 (d, J = 3.0 Hz), 117.5, 116.0 (d, J = 19.8 Hz), 116.0, 111.8, 111.6, 104.3, 97.9, 58.6, 55.7, 51.9, 20.9.

¹⁹**F NMR** (376 MHz, CDCl₃) δ -139.8 (s, 1F).

HRMS (ESI) m/z calcd. for C₂₅H₂₂O₂NCl₃F [M+H]⁺ 492.06947, found 492.06964.

General procedure C: Cu/CPA-catalyzed asymmetric three-component radical-initiated trifluoromethylarylation of 1,1-diarylalkene



Under argon, an oven-dried resealable Schlenk tube equipped with a magnetic stir bar was charged with 1,1-diarylalkene substrate 1 (0.1 mmol, 1.0 equiv.), substituted indole (0.12 mmol, 1.2 equiv.), CuI (3.0 mg, 0.015 mmol, 15 mol%), chiral phosphoric acid ((S)-A5 (9.86 mg, 0.01 mmol, 10 mol%), 3d (39.6 mg, 0.12 mmol, 1.2 equiv.), and anhydrous CHCl₃ (1.0 mL) at 0 °C, then the sealed tube was stirred at 0 °C for 40-72 h. Upon completion (monitored by TLC), the reaction mixture was directly purified by a silica gel chromatography [eluent: *n*-hexane/ethyl acetate = 100/0 to 2/1, using *n*-hexane (100%) to remove the solvent (DCM) at first] to afford the desired product 7.

Note: Since the reaction is sensitive to water and air, Schlenk tube and the reagents must be dried prior to use.



The racemate was prepared following the same procedure described above using CuI (3.0 mg, 0.015 mmol, 15 mol%) and diphenyl phosphate (2.5 mg, 0.01 mmol, 10 mol%) as catalyst at -35 °C in anhydrous DCM (1.0 mL) for 48 h. Upon completion (monitored by TLC), the solvent was removed *in vacuo*, and the residue was purified by a silica gel column chromatography (eluent: petroleum ether/EtOAc = 20/0-2/1) to give the desired product.

Note: Since the reaction is sensitive to water and air, Schlenk tube and the reagents must be dried prior to use.



(S)-4-(1-(5-bromo-1*H*-indol-3-yl)-3,3,3-trifluoro-1-(*p*-tolyl)p ropyl)phenol (7A)

The reaction was conducted on a 0.1 mmol scale according to the above general procedure C. The product was purified by silica gel flash column chromatography (n-hexane/ethyl acetate

= 100/0 to 2/1) to afford desired product 7A (35.3 mg, 75 %) as sticky light yellow oil.

HPLC analysis: Chiralcel AD3 (*n*-hexane/*i*-PrOH = 85/15, flow rate 0.5 mL/min, $\lambda = 240$ nm), t_R (major) = 31.92 min, t_R (minor) = 35.09 min.

¹**H NMR** (500 MHz, CDCl₃) δ 8.12 (s, 1H), 7.33-7.17 (m, 6H), 7.15-7.04 (m, 3H), 7.00 (s, 1H), 6.73 (d, J = 8.2 Hz, 2H), 4.91 (br s, 1H), 3.68-3.43 (m, 2H), 2.32 (s, 3H). ¹³**C NMR** (126 MHz, CDCl₃) δ 153.9, 141.8, 136.9, 136.1, 135.4, 129.6, 128.8, 128.1, 127.9, 125.9 (q, J = 280.0 Hz), 125.5, 124.8, 124.2, 120.4, 114.8, 112.7, 112.6, 48.7, 43.7 (q, J = 26.3 Hz), 20.9.

¹⁹F NMR (376 MHz, CDCl₃) δ -56.5 (t, J = 10.6 Hz, 3F). HRMS (ESI) m/z calcd. for C₂₄H₁₈ONBrF₃ [M-H]⁻ 472.05293, found 472.05203.



(S)-4-(3,3,3-trifluoro-1-(1*H*-indol-3-yl)-1-(*p*-tolyl)propyl)phen ol (7B)

The reaction was conducted on a 0.1 mmol scale according to the above general procedure C. The product was purified by silica gel flash column chromatography (n-hexane/ethyl acetate =

100/0 to 2/1) to afford desired product **7B** (37.9 mg, 96 %) as sticky light yellow oil. **HPLC** analysis: Chiralcel AD3 (*n*-hexane/*i*-PrOH = 85/15, flow rate 0.5 mL/min, λ = 230 nm), *t*_R (minor) = 22.02 min, *t*_R (major) = 25.13 min.

¹**H NMR** (500 MHz, CDCl₃) δ 8.03 (s, 1H), 7.41-7.19 (m, 5H), 7.19-7.00 (m, 4H), 6.99-6.86 (m, 2H), 6.69 (d, *J* = 7.7 Hz, 2H), 3.77 (br s, 1H), 3.60 (d, *J* = 10.2 Hz, 2H), 2.31 (s, 3H).

¹³**C NMR** (126 MHz, CDCl₃) δ 153.8, 142.2, 137.2, 136.8, 135.9, 129.8, 128.6, 128.1, 126.3, 126.1 (q, *J* = 280.0 Hz), 124.2, 121.9, 121.8, 120.8, 119.2, 114.7, 111.3, 48.8, 44.0, 43.7 (q, *J* = 26.0 Hz), 20.9.

¹⁹**F NMR** (376 MHz, CDCl₃) δ -56.4 (s, 3F).

HRMS (ESI) m/z calcd. for C₂₄H₁₉ONF₃ [M-H]⁻ 394.14242, found 394.14157.



(S)-4-(3,3,3-trifluoro-1-(5-methoxy-1*H*-indol-3-yl)-1-(*p*-toly l)propyl)phenol (7C)

The reaction was conducted on a 0.1 mmol scale according to the above general procedure C. The product was purified by silica gel flash column chromatography (*n*-hexane/ethyl acetate

= 100/0 to 2/1) to afford desired product 7C (40.0 mg, 94 %) as sticky light yellow oil. HPLC analysis: Chiralcel AD3 (*n*-hexane/*i*-PrOH = 85/15, flow rate 0.5 mL/min, λ = 240 nm), t_R (major) = 31.92 min, t_R (minor) = 35.09 min.

¹**H** NMR (500 MHz, CDCl₃) δ 7.99 (s, 1H), 7.36-7.21 (m, 6H), 7.09 (d, J = 8.1 Hz, 2H), 7.01 (d, J = 2.2 Hz, 1H), 6.81 (dd, J = 8.8, 2.2 Hz, 1H), 6.72 (d, J = 8.7 Hz, 2H), 6.39 (d, J = 1.9 Hz, 1H), 3.65-3.51 (m, 5H), 2.32 (s, 3H).

¹³**C NMR** (126 MHz, CDCl₃) δ 153.9, 153.2, 142.1, 137.0, 135.8, 132.0, 129.8, 128.6, 128.1, 126.9, 126.1 (q, *J* = 280.0 Hz), 124.9, 120.5, 114.7, 111.8, 111.6, 104.3, 55.7, 48.8, 43.8 (q, *J* = 26.1 Hz), 20.9.

¹⁹**F NMR** (376 MHz, CDCl₃) δ -56.4 (s, 3F).

HRMS (ESI) m/z calcd. for C₂₅H₂₁O₂NF₃ [M-H]⁻ 424.15299, found 424.15247.



(S)-4-(3,3,3-trifluoro-1-(5-fluoro-1*H*-indol-3-yl)-1-(*p*-tolyl)pr opyl)phenol (7D)

The reaction was conducted on a 0.1 mmol scale according to the above general procedure C. The product was purified by silica gel flash column chromatography (n-hexane/ethyl acetate =

100/0 to 2/1) to afford desired product **7D** (35.9 mg, 87 %) as sticky light yellow oil. **HPLC** analysis: Chiralcel AD3 (*n*-hexane/*i*-PrOH = 85/15, flow rate 0.5 mL/min, λ = 230 nm), *t*_R (minor) = 19.47 min, *t*_R (major) = 24.01 min.

¹**H NMR** (500 MHz, CDCl₃) δ 8.08 (s, 1H), 7.34-7.18 (m, 5H), 7.15-7.02 (m, 3H), 6.89 (t, J = 8.8 Hz, 1H), 6.74 (d, J = 8.1 Hz, 2H), 6.62 (d, J = 10.3 Hz, 1H), 4.77 (s, 1H), 3.55 (q, J = 10.5 Hz, 2H), 2.32 (s, 3H).

¹³C NMR (126 MHz, CDCl₃) δ 157.1 (d, J = 234.0 Hz), 153.9, 141.8, 136.9, 136.0, 133.3, 129.6, 128.7, 127.9, 126.8 (d, J = 10.0 Hz), 125.9 (q, J = 279.5 Hz), 125.88, 120.9 (d, J = 4.5 Hz), 114.8, 111.8 (d, J = 9.8 Hz), 110.3 (d, J = 26.4 Hz), 106.8 (d, J = 24.3 Hz), 48.6, 43.6 (q, J = 26.3 Hz), 20.9.

¹⁹F NMR (376 MHz, CDCl₃) δ -56.6 (s, 3F), -124.0 (s, 1F).

HRMS (ESI) m/z calcd. for C₂₄H₁₈ONF₄ [M-H]⁻ 412.13190, found 412.13217.



(S)-4-(1-(4-(*tert*-butyl)phenyl)-3,3,3-trifluoro-1-(5-methoxy-1*H*-indol-3-yl)propyl)phenol (7E)

The reaction was conducted on a 0.1 mmol scale according to the above general procedure C. The product was purified by silica gel flash column chromatography (n-hexane/ethyl acetate

= 100/0 to 2/1) to afford desired product **7E** (40.1 mg, 86 %) as sticky light yellow oil. **HPLC** analysis: Chiralcel OD3 (*n*-hexane/*i*-PrOH = 85/15, flow rate 0.5 mL/min, λ = 230 nm), *t*_R (major) = 14.57 min, *t*_R (minor) = 18.80 min.

¹**H NMR** (500 MHz, CDCl₃) δ 7.97 (s, 1H), 7.38-7.18 (m, 7H), 7.00 (s, 1H), 6.79 (d, *J* = 8.6 Hz, 1H), 6.72 (d, *J* = 8.4 Hz, 2H), 6.33 (s, 1H), 3.56 (s, 5H), 1.29 (s, 9H).

¹³**C NMR** (126 MHz, CDCl₃) δ 153.8, 153.2, 148.9, 142.2, 136.9, 131.9, 129.8, 127.7, 126.9, 126.1 (q, *J* = 280.2 Hz), 125.0, 124.8, 120.6, 114.6, 111.7, 104.2, 55.7, 48.6, 43.7 (q, *J* = 26.1 Hz), 34.3, 31.3.

¹⁹F NMR (376 MHz, CDCl₃) δ -56.3 (s, 3F).

HRMS (ESI) m/z calcd. for C₂₈H₂₉O₂NF₃ [M+H]⁺ 468.2150, found 468.2151.



(S)-4-(3,3,3-trifluoro-1-(4-hydroxyphenyl)-1-(5-methoxy-1*H* -indol-3-yl)propyl)benzaldehyde (7F)

The reaction was conducted on a 0.1 mmol scale according to the above general procedure C. The product was purified by silica gel flash column chromatography (n-hexane/ethyl acetate

= 100/0 to 2/1) to afford desired product **7F** (26.7 mg, 61 %) as sticky light yellow oil. **HPLC** analysis: Chiralcel IC (*n*-hexane/*i*-PrOH = 85/15, flow rate 0.3 mL/min, λ = 254 nm), t_R (minor) = 30.32 min, t_R (major) = 33.10 min. ¹**H** NMR (500 MHz, CDCl₃) δ 9.93 (s, 1H), 8.15 (s, 1H), 7.76 (d, *J* = 7.9 Hz, 2H), 7.59 (d, *J* = 7.9 Hz, 2H), 7.23 (dd, *J* = 16.5, 7.7 Hz, 3H), 6.99 (s, 1H), 6.80 (d, *J* = 8.8 Hz, 1H), 6.72 (d, *J* = 8.2 Hz, 2H), 6.32 (s, 1H), 5.29 (s, 1H), 3.64-3.49 (m, 5H).

¹³**C NMR** (126 MHz, CDCl₃) δ 192.3, 154.4, 153.3, 152.0, 135.5, 134.4, 132.0, 129.7, 129.5, 128.9, 126.5, 125.8 (q, *J* = 279.5 Hz), 125.0, 119.2, 115.0, 112.0, 111.7, 104.1, 55.8, 49.5, 43.5 (q, *J* = 26.6 Hz).

¹⁹**F NMR** (376 MHz, CDCl₃) δ -56.5 (s, 3F).

HRMS (ESI) m/z calcd. for C₂₅H₁₉O₃NF₃ [M+H]⁺ 438.13115, found 438.13126.



(S)-4-(3,3,3-trifluoro-1-(5-methoxy-1*H*-indol-3-yl)-1-(4-nitro phenyl)propyl)phenol (7G)

The reaction was conducted on a 0.1 mmol scale according to the above general procedure C. The product was purified by silica gel flash column chromatography (n-hexane/ethyl acetate

= 100/0 to 2/1) to afford desired product 7G (27.7 mg, 61 %) as a sticky light yellow solid.

HPLC analysis: Chiralcel AD3 (*n*-hexane/*i*-PrOH = 80/20, flow rate 0.7 mL/min, λ = 214 nm), t_R (minor) = 24.54 min, t_R (major) = 29.43 min.

¹**H** NMR (500 MHz, CDCl₃) δ 8.20 (s, 1H), 8.11 (d, J = 8.5 Hz, 2H), 7.60 (d, J = 8.5 Hz, 2H), 7.27 (d, J = 8.5 Hz, 1H), 7.21 (d, J = 8.2 Hz, 2H), 7.02 (s, 1H), 6.83 (d, J = 8.7 Hz, 1H), 6.74 (d, J = 8.2 Hz, 2H), 6.32 (s, 1H), 5.45 (br s, 1H), 3.67-3.52 (m, 5H). ¹³C NMR (126 MHz, CDCl₃) δ 154.6, 153.4, 152.3, 146.2, 135.1, 132.1, 129.7, 129.2, 126.3, 125.7 (q, J = 280.0 Hz), 124.9, 123.2, 118.9, 115.1, 112.2, 111.8, 104.0, 55.8, 49.4 (d, J = 1.4 Hz), 43.6 (q, J = 26.6 Hz).

¹⁹**F NMR** (376 MHz, CDCl₃) δ -56.5 (s, 3F).

HRMS (ESI) m/z calcd. for C₂₄H₁₈O₄N₂F₃ [M+H]⁺ 455.12241, found 455.12161.



(*S*)-4-(3,3,3-trifluoro-1-(5-methyl-1*H*-pyrrol-2-yl)-1-(*p*-tolyl) propyl)phenol (7H)

The reaction was conducted on a 0.1 mmol scale according to the above general procedure C. The product was purified by silica gel flash column chromatography (n-hexane/ethyl acetate

= 100/0 to 2/1) to afford desired product 7A (28.0 mg, 78 %) as sticky light yellow oil. HPLC analysis: Chiralcel OD3 (*n*-hexane/*i*-PrOH = 85/15, flow rate 0.3 mL/min, λ = 230 nm), *t*_R (major) = 18.70 min, *t*_R (minor) = 20.94 min.

¹**H** NMR (500 MHz, CDCl₃) δ 7.36 (s, 1H), 7.13 (s, 4H), 6.97 (d, J = 8.5 Hz, 2H), 6.77 (d, J = 9.0 Hz, 2H), 5.94 (t, J = 3.0 Hz, 1H), 5.82-5.79 (m, 1H), 5.10 (br s, 1H), 3.40 (q, J = 10.5 Hz, 2H), 2.36 (s, 3H), 2.17 (s, 3H).

¹³**C NMR** (125 MHz, CDCl₃) δ 154.3, 141.6, 136.5, 136.4, 133.9, 129.9, 128.9, 128.2, 127.8, 126.3 (q, *J* = 277.3 Hz), 114.8, 107.6, 105.3, 49.29, 44.2 (q, *J* = 26.5 Hz), 20.9, 13.1.

¹⁹**F NMR** (376 MHz, CDCl₃) δ -56.1 (s, 3F).

HRMS (ESI) m/z calcd. for $C_{21}H_{21}F_3NO [M+H]^+$ 360.1570, found 360.1565.
Gram-scale reaction with diverse radical precursors



Synthesis of 4C:

The reaction was conducted on a 4.0 mmol (1.008 g) scale according to the above general procedure A. The product was purified by silica gel flash column chromatography (*n*-hexane/ethyl acetate = 100/0 to 2/1) to afford desired product 4C (2.314 g, 87 %) as sticky light yellow oil.

Synthesis of 6E:



(S)-4-(1-(4-(*tert*-butyl)phenyl)-3,3,3-trichloro-1-(5-methoxy -1H-indol-3-yl)propyl)phenol (6E)

The reaction was conducted on a 4.0 mmol (1.008 g) scale according to the above general procedure B. The product was purified by silica gel flash column chromatography

(*n*-hexane/ethyl acetate = 100/0 to 2/1) to afford desired product **6E** (1.648 mg, 80 %) as sticky light yellow oil.

Note: the pure **6E** is unstable at 25-35°C, easy to convert to dichloroolefin-containing product or other products, especially when dissolved in halogen-containing solvents under air, such as DCM, CHCl₃ etc. **6E** gradually converted to dichloroolefin-containing product or other products in other solvents, such as n-hexane, ethyl acetate, alcohol isopropanol etc at 25-30°C under air atmosphere. So **6E** should be characterized by NMR, HPLC or HRMS ASAP.

As the pure **6E** is unstable at 20-35°C, amounts of EtOAc is observed in NMRs.

HPLC analysis: Chiralcel IA (*n*-hexane/*i*-PrOH = 90/10, flow rate 0.4 mL/min, $\lambda = 260$ nm), t_R (major) = 33.72 min, t_R (minor) = 37.56 min.

¹**H** NMR (400 MHz, CDCl₃) δ 8.09 (s, 1H), 7.49 (d, J = 8.2 Hz, 4H), 7.24 (dd, J = 18.6, 8.3 Hz, 4H), 6.78 (d, J = 8.7 Hz, 1H), 6.72 (d, J = 8.1 Hz, 2H), 6.21 (s, 1H), 4.42-4.24 (m, 2H), 3.54 (s, 3H), 1.28 (s, 9H).

¹³C NMR (101 MHz, CDCl₃) δ 153.8, 153.1, 148.7, 142.9, 137.2, 131.7, 129.8, 127.8, 127.5, 126.7, 124.8, 118.0, 114.7, 111.7, 104.2, 98.2, 58.7, 55.5, 51.7, 34.3, 31.3. HRMS (ESI) m/z calcd. for C₂₈H₂₉O₂NCl₃ [M+H]⁺516.12584, found 516.12630.

Synthesis of **7E**: The reaction was conducted on a 4.0 mmol (1.008 g) scale according to the above general procedure C. The product was purified by silica gel flash column chromatography (*n*-hexane/ethyl acetate = 100/0 to 2/1) to afford desired product **7E** (1.663 g, 89 %) as sticky light yellow oil.

Procedure for synthetic application:



Synthesis of 7Ba: 7Ba was synthesized according to the procedures previously reported.⁴



(S)-4-(3,3,3-trifluoro-1-(1*H*-indol-3-yl)-1-(*p*-tolyl)propyl)phe nyl trifluoromethanesulfonate (7Ba)

HPLC analysis: Chiralcel OD-H (*n*-hexane/*i*-PrOH = 90/10, flow rate 0.8 mL/min, $\lambda = 254$ nm), $t_{\rm R}$ (minor) = 10.93 min, $t_{\rm R}$ (major) = 18.24 min.

¹**H** NMR (500 MHz, CDCl₃) δ 8.09 (s, 1H), 7.48 (d, *J* = 8.9 Hz, 2H), 7.34 (d, *J* = 8.2 Hz, 1H), 7.28-7.21 (m, 2H), 7.14 (t, *J* = 8.0 Hz, 3H), 7.07 (d, *J* = 8.1 Hz, 2H), 6.98-6.87 (m, 3H), 3.59 (q, *J* = 10.5 Hz, 2H), 2.30 (s, 3H).

¹³**C NMR** (126 MHz, CDCl₃) δ 147.9, 145.4, 141.0, 136.9, 136.5, 130.5, 128.9, 128.1, 126.0, 125.8 (q, *J* = 280.0 Hz), 124.2, 122.1, 121.6, 120.6, 118.7 (q, *J* = 321.3 Hz), 119.9, 119.5, 111.5, 49.3 (d, *J* = 1.5 Hz), 43.7 (q, *J* = 26.6 Hz), 20.9.

¹⁹F NMR (376 MHz, CDCl₃) δ -56.4 (s, 3F), -72.9 (s, 3F).

HRMS (ESI) m/z calcd. for C₂₅H₁₈O₃NF₆S [M-H]⁻ 526.09171, found 526.09045.

Synthesis of 7Bb: 7Bb was synthesized according to the procedures previously reported.⁵



(S)-3-(3,3,3-trifluoro-1-phenyl-1-(*p*-tolyl)propyl)-1*H*-indole (7Bb)

HPLC analysis: Chiralcel OD3 (*n*-hexane/*i*-PrOH = 85/15, flow rate 0.6 mL/min, $\lambda = 254$ nm), $t_{\rm R}$ (minor) = 9.98 min, $t_{\rm R}$ (major) = 10.57 min.

¹**H NMR** (500 MHz, CDCl₃) δ 8.03 (s, 1H), 7.40 (d, J = 7.7 Hz, 2H), 7.33 (d, J = 8.2 Hz, 1H), 7.31-7.21 (m, 4H), 7.18 (t, J = 7.2 Hz, 1H), 7.12 (t, J = 7.6 Hz, 1H), 7.06 (d, J = 8.1 Hz, 2H), 7.01-6.94 (m, 2H), 6.89 (t, J = 7.5 Hz, 1H), 3.62 (q, J = 10.7 Hz, 2H), 2.29 (s, 3H).

¹³C NMR (126 MHz, CDCl₃) δ 145.0, 141.9, 136.8, 135.9, 128.7, 128.3, 128.2, 127.9, 126.40, 126.35, 126.0 (q, J = 279.9 Hz), 124.4, 121.9, 121.8, 120.5, 119.2, 111.3, 49.5 (d, J = 1.6 Hz), 43.5 (q, J = 26.3 Hz), 20.9. ¹⁹F NMR (376 MHz, CDCl₃) δ -56.5 (s, 3F). HRMS (ESI) m/z calcd. for C₂₄H₁₉NF₆ [M-H]⁻ 378.14751, found 378.14661.

Synthesis of **7Bc**: **7Bc** was synthesized according to the procedures previously reported.⁶



(S)-3-(3,3,3-trifluoro-1-(4'-methoxy-[1,1'-biphenyl]-4-yl)-1-(*p*-tolyl)propyl)-1*H*-indole (7Bc) HPLC analysis: Chiralcel AZ3 (*n*-hexane/*i*-PrOH = 90/10, flow rate 0.4 mL/min, $\lambda = 254$ nm), t_R (minor) = 19.31 min, t_R (major) = 21.19 min.

¹**H** NMR (500 MHz, CDCl₃) δ 8.05 (s, 1H), 7.50 (t, J = 5.7 Hz, 2H), 7.43 (s, 4H), 7.32 (dd, J = 10.6, 8.3 Hz, 3H), 7.13 (t, J = 7.6 Hz, 1H), 7.07 (d, J = 8.2 Hz, 2H), 7.01 (dd, J = 14.0, 5.3 Hz, 2H), 6.97-6.87 (m, 3H), 3.82 (s, 3H), 3.64 (q, J = 10.7 Hz, 2H), 2.30 (s, 3H).

¹³C NMR (126 MHz, CDCl₃) δ 159.0, 143.5, 141.9, 138.4, 136.8, 135.9, 133.1, 128.7, 128.6, 128.2, 127.9, 126.4, 126.03 (q, J = 277.3 Hz), 126.01, 124.5, 121.9, 121.8, 120.3, 119.2, 114.1, 111.3, 55.3, 49.2 (d, J = 1.6 Hz), 43.4 (q, J = 26.3 Hz), 20.9. ¹⁹F NMR (376 MHz, CDCl₃) δ -56.5 (s, 3F).

HRMS (ESI) m/z calcd. for C₃₁H₂₅ONF₃ [M-H]⁻ 484.18937, found 484.18845.

Synthesis of 8A: 8A was synthesized according to the procedures previously reported.⁷



(S)-4-(1-(5-bromo-1*H*-indol-3-yl)-3,3-difluoro-4-hydroxy-1-(*p*-tolyl)butyl)phenol (8A)

HPLC analysis: Chiralcel AD3 (*n*-hexane/*i*-PrOH = 75/25, flow rate 0.6 mL/min, $\lambda = 254$ nm), $t_{\rm R}$ (minor) = 17.43 min, $t_{\rm R}$ (major) = 23.25 min.

¹**H NMR** (500 MHz, CDCl₃) δ 8.11 (s, 1H), 7.35-7.29 (m, 4H), 7.26-7.18 (m, 2H), 7.15 (d, J = 5.2 Hz, 2H), 7.08 (d, J = 8.1 Hz, 2H), 6.74 (d, J = 8.6 Hz, 2H), 4.74 (s, 1H), 3.41 (td, J = 16.4, 3.9 Hz, 2H), 3.11 (t, J = 13.3 Hz, 2H), 2.32 (s, 3H).

¹³**C NMR** (126 MHz, CDCl₃) δ 152.7, 141.6, 136.8, 134.8, 134.3, 128.8, 127.6, 127.2, 127.1, 124.4, 123.7, 123.3, 120.5, 113.6, 111.7, 111.5, 64.0 (t, *J* = 31.9 Hz), 48.1, 41.5 (t, *J* = 22.7 Hz), 28.7, 19.9.

¹⁹**F NMR** (376 MHz, CDCl₃) δ -97.1 (s, 2F). **UDMS** (ESI) m/z colled for Couller On NBrEe [M 11]² 484.07202 four

HRMS (ESI) m/z calcd. for $C_{25}H_{21}O_2NBrF_2$ [M-H]⁻ 484.07292, found 484.07196.

Synthesis of **8B**: **8B** was synthesized according to the procedures previously reported with minor revision.⁷ NaH (8.0 mg, 60% in mineral oil, 0.2 mmol) was added to a stirred solution of **5A** (51.5 mg, 0.1 mmol) in dry THF (2.0 mL) in an oven-dried Schlenk tube at rt. After stirring at rt for additional 2 h, the reaction mixture was quenched with saturated NH4Cl solution and stirred for 15 min. The mixture was extracted three times with ethyl acetate, dried over Na2SO4, and concentrated *in vacuo*.

The crude product was purified by flash column chromatography on silica gel (eluent: EtOAc/MeOH = 100/1 to 20/1) to give **8B** (37.4 mg, 75%).



(S)-4-(5-bromo-1*H*-indol-3-yl)-2,2-difluoro-4-(4-hydroxyphe nyl)-4-(*p*-tolyl)butanoic acid (8B)

¹**H** NMR (500 MHz, MeOD) δ 7.28 (d, J = 8.1 Hz, 2H), 7.22 (dd, J = 8.6, 4.1 Hz, 3H), 7.16 (s, 1H), 7.07 (d, J = 8.6 Hz, 1H), 7.02 (d, J = 8.0 Hz, 2H), 6.97 (d, J = 1.6 Hz, 1H), 6.66 (d, J =

8.7 Hz, 2H), 3.53 (t, *J* = 17.6 Hz, 2H), 3.37 (s, 1H), 2.28 (s, 3H). ¹³C NMR (126 MHz, MeOD) δ 171.4, 154.9, 143.5, 137.0, 135.8, 134.9, 129.8, 128.5, 128.4, 127.7, 126.0, 123.8, 123.0, 121.1, 113.7, 112.4, 110.9, 49.1, 48.5, 44.0 (t, *J* = 22.4 Hz), 19.6.

¹⁹**F NMR** (376 MHz, MeOD) δ -96.6 (s, 2F).

HRMS (ESI) m/z calcd. for C₂₅H₁₉O₃NBrF₂ [M-H]⁻ 498.05219, found 498.05182.



(S)-4-(3,3-dichloro-1-(5-methoxy-1*H*-indol-3-yl)-1-(*p*-tolyl)al lyl)phenol (9A)

The reaction was conducted on a 0.1 mmol scale according to the above general procedure B. The product was purified by silica gel flash column chromatography (n-hexane/ethyl acetate

= 100/0 to 2/1) to afford desired trichloromethyl-product, which gradually converted to dichloroolefin-containing product **9A** (39.3 mg, 90 %) as sticky black oil when concentrated at 40°C *in vacuo*.

Note: the pure **9A** is unstable at 25-35°C, easy to convert to other products, especially when dissolved in halogen-containing solvents under air, such as DCM, CHCl₃ etc. So **9A** should be characterized by NMR, HPLC or HRMS ASAP.

HPLC analysis: Chiralcel AD3 (*n*-hexane/*i*-PrOH = 85/15, flow rate 0.6 mL/min, λ = 230 nm), *t*_R (minor) = 17.17 min, *t*_R (major) = 23.97 min.

¹**H NMR** (500 MHz, CDCl₃) δ 7.93 (s, 1H), 7.23 (d, J = 8.8 Hz, 1H), 7.12-7.06 (m, 7H), 7.00 (s, 1H), 6.81 (dd, J = 8.8, 2.3 Hz, 1H), 6.73 (d, J = 8.6 Hz, 2H), 6.66 (d, J = 1.9 Hz, 1H), 6.43 (d, J = 2.2 Hz, 1H), 3.61 (s, 3H), 2.32 (s, 3H).

¹³C NMR (126 MHz, CDCl₃) δ 154.1, 153.5, 141.5, 136.7, 136.6, 136.1, 132.2, 130.6, 129.1, 128.7, 126.8, 126.0, 123.0, 121.2, 114.8, 112.0, 111.8, 103.4, 55.8, 54.9, 21.0. HRMS (ESI) m/z calcd. for C₂₅H₂₂O₂NCl₂ [M+H]⁺ 438.10221, found 438.10245.

Mechanistic study



a) Trapping with TEMPO



Under argon atmosphere, an oven-dried resealable Schlenk tube equipped with a magnetic stir bar was charged with 1,1-diarylalkene **1a** (0.05 mmol, 1.0 equiv.), 5-bromo-1*H*-indole (0.06 mmol, 1.2 equiv.), CuI (1.0 mg, 0.005 mmol, 10 mol%), chiral phosphoric acid ((*S*)-**A5** (5 mg, 0.005 mmol, 10 mol%), *n*-C4F9SO₂Cl (**3a**, 19.1 mg, 0.06 mmol, 1.2 equiv.), 2,2,6,6-tetramethyl-1-piperidinyloxy (TEMPO, 9.4 mg, 0.06 mmol, 1.2 equiv), Ag₂CO₃ (8.3 mg, 0.03 mmol, 0.6 equiv.), and anhydrous DCM (0.5 mL) at 0 °C, then the sealed tube was stirred at 0 °C for 40 h. Conversion was based on ¹H NMR/¹⁹F NMR/GC-MS analysis of the crude product.

Note: Since the reaction is sensitive to water and air, Schlenk tube and the reagents must be dried prior to use.





b) Linear effect experiments



These reactions were conducted according to the general procedure A. The product was separated by preparative TLC. Chiral HPLC analysis gave linear effect data. The ee of (*R*)-4A was determined by HPLC analysis: Chiralcel IA (*n*-hexane/*i*-PrOH = 85/15, flow rate 0.5 mL/min, $\lambda = 240$ nm), t_R (major) = ~10.57 min, t_R (minor) = ~11.72 min.

Supplementary references:

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NMR spectra





$\begin{array}{c} 7.284 \\ 6.591 \\ 6.594 \\ 6.594 \\ 6.594 \\ 6.594 \\ 6.594 \\ 6.592 \\ 5.572 \\ 5.2272 \\ 5.2272 \\ 5.2272 \\ 5.2272 \\ 5.2272 \\ 5.2233 \\ 5.2233 \\ 5.233 \\ 5.233$











S51







5Ba







S56











S61

















-8.164 -7.570 -7.518 -7.518 -7.518 -7.518 -7.518 -7.751 -7.752 -7.752 -7.7577 -7.7577 -7.7577 -7.7577 -7.7577 -7.7577 -7.7577 -7.7577 -7.7577


























 $\begin{array}{c} & & 8 \\$































S96







S99










































HPLC spectra













Peak #	RetTime [min]	Туре	Width [min]	Area [mAU*s]	Height [mAU]	Area %
1	19.084	VB	0.3582	7664.07861	328.44595	49.9814
2	22.037	MF	0.4779	7669.79102	267.47556	50.0186
Total	ls :			1.53339e4	595.92151	









Peak #	RetTime [min]	Туре	Width [min]	Area [mAU*s]	Height [mAU]	Area %
1	25.111	BB	0.6511	1547.42676	36.46709	50.0593
2	27.909	BB	0.7841	1543.76282	29.91044	49.9407
Total	s :			3091.18958	66.37753	





S127





Totals : 1.53901e4 403.95433





S131



S132







S135























Peak #	RetTime [min]	Туре	Width [min]	Area [mAU*s]	Height [mAU]	Area %
1	33.721	BB	0.8365	2514.59985	44.27938	92.9835
2	37.539	BB	0.6441	189.75143	3.45831	7.0165
Totals :				2704.35129	47.73769	


S145



Signal 1: DAD1 A, Sig=254,4 Ref=360,100

1 9.975 MM 0.2141 15.31647 1.19216 2.0208	Peak #	RetTime [min]	Туре	Width [min]	Area [mAU*s]	Height [mAU]	Area %
	1	9.975	MM	0.2141	15.31647	1.19216	2.0208



Signal 2: DAD1 B, Sig=254,4 Ref=360,100

Peak #	RetTime [min]	Туре	Width [min]	Area [mAU*s]	Height [mAU]	Area %
1	19.457	BB	0.4756	2765.72485	89.71932	50.0864
2	21.454	BB	0.4799	2756.17969	88.35643	49.9136

Totals :

5521.90454 178.07575







Totals :	1.19409e4	247.06272