Supporting Information

for

One-pot gold-catalyzed synthesis of 3-silylethynyl indoles

from unprotected o-alkynylanilines

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General procedures

All reactions were carried out in oven dried glassware under an atmosphere of nitrogen, unless stated otherwise. For quantitative flash chromatography technical grade solvents were used. For flash chromatography for analysis, HPLC grade solvents from Sigma-Aldrich were used. THF, Et₂O, CH₃CN, toluene, hexane and CH₂Cl₂ were dried by passage over activated alumina under a nitrogen atmosphere (H₂O content < 10 ppm, Karl-Fischer titration). NaAuCl₄ and AuCl were purchased from Aldrich and stored in desiccators under anhydrous condition [decrease of reactivity has been observed for the catalysts on prolonged exposed to air (ca. 1 month)]. All chemicals were purchased from Acros, Aldrich, Fluka, VWR, Aplichem, Maybridge, TCI or Merck and used as received unless stated otherwise. Chromatographic purification was performed by flash chromatography on Macherey-Nagel silica 40–63, 60 Å, using the solvents indicated as eluent at 0.1–0.5 bar pressure. TLC was performed on Merck silica gel 60 F₂₅₄ TLC glass plates or aluminium plates and visualized with UV light, permanganate stain, CAN stain or anisaldehyde stain. Melting points were measured on a Büchi B-540 melting point apparatus using open glass capillaries, the data is corrected. ¹H NMR spectra were recorded on a Bruker DPX-400 400 MHz spectrometer in CDCl₃, DMSO-d₆ or CD₃OD, all signals are reported in ppm with the internal chloroform signal at 7.26 ppm, the internal DMSO signal at 2.50 ppm or the internal methanol signal at 3.30 ppm as standard. The data is reported as (s = singlet, d = doublet, t = triplet, q = quadruplet, qi = quintet, m = multiplet or unresolved, br = broad signal, app = apparent coupling constant(s) in Hz, integration, interpretation). ¹³C NMR spectra were recorded with ¹H-decoupling on a Bruker DPX-400 spectrometer at 100 MHz in CDCl₃, DMSO-d₆ or CD₃OD, all signals are reported in ppm with the internal chloroform signal at 77.0 ppm, the internal DMSO signal at 39.5 ppm or the internal methanol signal at 49.0 ppm as standard. Infrared spectra were recorded on a JASCO FT-IR B4100 spectrophotometer with an ATR PRO410-S and a ZnSe prism and are reported in cm⁻¹ (w = weak, m = medium, s = strong, br = broad). Gas chromatographic and low resolution mass spectrometric measurements were performed on a Perkin-Elmer Clarus 600 gas chromatography and mass spectrometer using a Perkin-Elmer Elite fused silica column (length: 30 m, diameter: 0.32 mm) and helium as the carrier gas. High resolution mass spectrometric measurements were performed by the mass spectrometry service of ISIC at the EPFL on a MICROMASS (ESI) Q-TOF Ultima API.
TIPS-EBX (1) synthesis

1-Hydroxy-1,2-benziodoxol-3(1H)-one (7)

Following a reported procedure [1], NaIO$_4$ (6.7 g, 31 mmol; 1.0 equiv) and 2-iodobenzoic acid (6) (7.4 g, 30 mmol, 1.0 equiv) were suspended in 30% (v:v) aq. AcOH (45 mL). The mixture was vigorously stirred and heated under reflux for 4 h. The reaction mixture was then diluted with cold water (120 mL) and allowed to cool to room temperature with protection from light. After 1 h, the crude product was collected by filtration, washed on the filter with ice cold water (3 × 30 mL) and acetone (3 × 30 mL), and air dried in the dark to give pure 7 (7.3 g, 19 mmol, 92% yield) as a colorless solid. $^1$H NMR (400 MHz, (CD$_3$)$_2$SO) δ 8.02 (dd, $J = 7.7$, 1.4 Hz, 1 H, ArH), 7.97 (m, 1 H, ArH), 7.85 (dd, $J = 8.2$, 0.7 Hz, 1 H, ArH), 7.71 (td, $J = 7.6$, 1.2 Hz, 1 H, ArH). $^{13}$C NMR (100 MHz, (CD$_3$)$_2$SO) δ 167.7, 134.5, 131.5, 131.1, 130.4, 126.3, 120.4. IR ν 3083 (w), 3060 (w), 2867 (w), 2402 (w), 1901 (m), 1585 (m), 1564 (m), 1440 (m), 1338 (s), 1302 (m), 1148 (m), 1018 (w), 834 (m), 798 (w), 740 (s), 694 (s), 674 (m), 649 (m). The characterization data for compounds 7 corresponded to the reported values [1].

Triisopropylsilyl trimethylsilylacetylene (9)

Following a reported procedure [2], n-butyllithium (2.5 M in hexanes, 12.0 mL, 29.9 mmol, 0.98 equiv) was added dropwise to a stirred solution of ethynyltrimethylsilane (8) (3.0 g, 30 mmol, 1.0 equiv) in THF (48 mL) at −78 °C. The mixture was then warmed to 0 °C and stirred for 5 min. The mixture was then re-cooled to −78 °C and chlorotriisopropylsilane (6.4 mL, 30 mmol, 1.0 equiv) added dropwise. The mixture was allowed to warm to room temperature and stirred overnight. A saturated solution of ammonium chloride (40 mL) was added, and the reaction mixture extracted with diethyl ether (2 × 60 mL). The organic layer was washed successively with water and brine, then dried over MgSO$_4$, filtered and concentrated under reduced pressure to give a colorless liquid which was further purified by Kugelrohr distillation (56–57 °C/0.25 mmHg) to yield 9 (7.16 g, 28.0 mmol, 92% yield) as a colorless liquid. $^1$H NMR (400 MHz, CDCl$_3$) δ 1.08 (m, 21 H, TIPS), 0.18 (s, 9 H, TMS). IR
Following a reported procedure [3], 2-iodosylbenzoic acid (7) (21.7 g, 82.0 mmol, 1.0 equiv) was charged in oven-dried three-neck 1L flask equipped with a magnetic stirrer. After 3 vacuum/nitrogen cycles, anhydrous acetonitrile (500 mL) was added via canula and cooled to 4 °C. Trimethylsilyltribrate (16.4 mL, 90.0 mmol, 1.1 equiv) was added dropwise via a dropping funnel over 30 min (no temperature increase was observed). After 15 min, (trimethylsilyl)(triisopropylsilyl)acetylene (9) (23.0 g, 90.0 mmol, 1.1 equiv) was added via canula over 15 min (no temperature increase was observed). After 30 min, the suspension became an orange solution. After 10 min, pyridine (7.0 mL, 90 mmol, 1.1 equiv) was added via syringe. After 15 min, the reaction mixture was transferred to a one-neck 1L flask and concentrated under vacuum until a solid was obtained. The solid was dissolved in DCM (200 mL) and transferred to a 1 L separatory funnel. The organic solution was added, washed with 1 M HCl (200 mL) and the aqueous layer extracted with CH$_2$Cl$_2$ (200 mL). The organic layers were combined, washed with a saturated solution of NaHCO$_3$ (2 × 200 mL), dried over MgSO$_4$, filtered and the solvent was evaporated under reduced pressure. Recrystallization from acetonitrile (ca. 120 mL) afforded 1 (30.1 g, 70.2 mmol, 86%) as colorless crystals. Mp (dec.) 170–176 °C. $^1$H NMR (400 MHz, CDCl$_3$) δ 8.44 (m, 1 H, ArH), 8.29 (m, 1 H, ArH), 7.77 (m, 2 H, ArH), 1.16 (m, 21 H, TIPS). $^{13}$C NMR (100 MHz, CDCl$_3$) δ 166.4, 134.6, 132.3, 131.4, 131.4, 126.1, 115.6, 114.1, 64.6, 18.4, 11.1. IR ν 2943 (m), 2865 (m), 1716 (m), 1618 (m), 1604 (s), 1584 (m), 1557 (m), 1465 (m), 1439 (w), 1349 (m), 1291 (m), 1270 (w), 1244 (m), 1140 (m), 1016 (m), 999 (m), 883 (m), 833 (m), 742 (m), 702 (s), 636 (m). Characterization data of 1 corresponded to the literature values [3].
2-Alkynylanilines synthesis

2-(Phenylethynyl)aniline (2a)

Following a slightly modified procedure [4], phenylacetylene (10) (1.20 ml, 11.0 mmol, 1.2 equiv) was added to a solution of 2-iodoaniline (4a) (2.0 g, 9.1 mmol, 1 equiv), PdCl₂(PPh₃)₂ (309 mg, 0.440 mmol, 0.05 equiv) and CuI (84 mg, 0.44 mmol, 0.2 equiv) in Et₃N (50 mL). The resulting suspension was stirred for 2 h under a nitrogen atmosphere at RT. The resulting mixture was filtered through Celite® and concentrated under vacuum. The resulting solid was purified by column chromatography (pentane/EtOAc 8/2) to afford 2a (1.85 g, 9.57 mmol, quant.) as an orange solid. Mp: 85–87 °C (lit [4] 91–92 °C). Rf (pentane/EtOAc 8/2): 0.4. ¹H NMR (400 MHz, CDCl₃) δ 7.54 (m, 2 H, ArH), 7.36 (m, 4 H, ArH), 7.15 (td, 1H, J = 7.7, 1.5 Hz, ArH), 6.73 (m, 2 H, ArH), 4.28 (s, 2 H, NH₂). ¹³C NMR (100 MHZ, CDCl₃) δ 147.7, 132.1, 131.4, 129.7, 128.4, 128.2, 123.3, 118.0, 114.3, 107.9, 94.7, 85.8. ¹H NMR is consistent with reported values [4].

2-(p-Tolylethynyl)aniline (2b)

Following a slightly modified procedure [5], a solution of 2-iodoaniline (4a) (329 mg, 1.50 mmol, 1 equiv), 1-ethynyl-4-methylbenzene (11) (209 mg, 1.80 mmol, 1.2 equiv), PdCl₂(PPh₃)₂ (102 mg, 0.150 mmol, 0.1 equiv) and CuI (28 mg, 0.15 mmol, 0.1 equiv) were heated under reflux in Et₃N (15 mL) for 2 h under a nitrogen atmosphere. The resulting mixture was filtered through Celite®, washed with DCM and concentrated under vacuum. The resulting solid was purified by column chromatography (pentane/EtOAc 95/5) to afford 2b (276 mg, 1.33 mmol, 88%) as a yellow solid. Mp: 104–107 °C. Rf (pentane/EtOAc 95/5): 0.3. ¹H NMR (400 MHz, CDCl₃) δ 7.45 (d, 2 H, J = 8.1 Hz, ArH), 7.38 (m, 1 H, ArH), 7.16 (m, 3 H, ArH), 6.74 (m, 2 H, ArH), 4.28 (s, 2 H, NH₂), 2.39 (s, 3 H, CH₃). ¹³C NMR (100 MHZ, CDCl₃) δ 147.7, 138.3, 132.0, 131.3, 129.5, 129.1, 120.2, 117.9, 114.2, 108.1, 94.8, 85.2, 21.5. IR ν 3475 (w), 3376 (w), 3056 (w), 3027 (w), 2919 (w), 2863 (w), 2207 (w), 1911 (w),
1611 (s), 1567 (w), 1509 (m), 1489 (m), 1456 (m), 1312 (m), 1258 (w), 1182 (w), 1157 (w),
1028 (w), 940 (w), 909 (w), 869 (w), 819 (s), 747 (s).

HRMS (ESI) calcd for C\textsubscript{15}H\textsubscript{14}N\textsuperscript{+} [M+H]\textsuperscript{+} 208.1121; found 208.1125.

2-((4-Fluorophenyl)ethynyl)aniline (2c)

Following a slightly modified procedure [5], a solution of 2-iodoaniline (4a) (329 mg, 1.50 mmol, 1 equiv), 1-ethynyl-4-fluorobenzene (12) (216 mg, 1.80 mmol, 1.2 equiv), PdCl\textsubscript{2}(PPh\textsubscript{3})\textsubscript{2} (105 mg, 0.15 mmol, 0.1 equiv) and CuI (28 mg, 0.15 mmol, 0.1 equiv) were heated under reflux in Et\textsubscript{3}N (15 mL) for 2 h under a nitrogen atmosphere. The resulting mixture was filtered through Celite\textsuperscript{®}, washed with DCM and concentrated under vacuum. The resulting solid was purified by column chromatography (pentane/EtOAc 8/1) to afford 2c (208 mg, 0.985 mmol, 66%) as a yellow solid. Mp: 97–98 °C (lit [5], 79–81 °C). Rf (pentane/EtOAc 8/1): 0.4. \textsuperscript{1}H NMR (400 MHz, CDCl\textsubscript{3}) \(\delta 7.50 (m, 2\ H, \text{ArH}), 7.35 (d, 1\ H, J = 8.1\ Hz, \text{ArH}), 7.15 (td, 1H, J = 7.9, 1.5 Hz, \text{ArH}), 7.05 (t, 2\ H, J = 8.7\ Hz, \text{ArH}), 6.72 (m, 2\ H, \text{ArH}), 4.25 (s, 2\ H, \text{NH})\). \textsuperscript{13}C NMR (100 MHZ, CDCl\textsubscript{3}) \(\delta 162.4 (d, J = 249\ Hz), 147.7, 133.2 (d, J = 8\ Hz), 132.0, 129.7, 119.3 (d, J = 3\ Hz), 117.9, 115.6 (d, J = 22\ Hz), 114.3, 107.6, 93.5, 85.5\). Consistent with reported values [5].

2-((4-Methoxyphenyl)ethynyl)aniline (2d)

Following a slightly modified procedure [5], a solution of 2-iodoaniline (4a) (329 mg, 1.50 mmol, 1 equiv), 1-ethynyl-4-methoxybenzene (13) (258 mg, 1.95 mmol, 1.2 equiv), PdCl\textsubscript{2}(PPh\textsubscript{3})\textsubscript{2} (105 mg, 0.15 mmol, 0.1 equiv) and CuI (28 mg, 0.15 mmol, 0.1 equiv) were heated under reflux in Et\textsubscript{3}N (15 mL) for 2 h under a nitrogen atmosphere. The resulting mixture was filtered through Celite\textsuperscript{®}, washed with DCM and concentrated under vacuum. The resulting solid was purified by column chromatography (pentane/EtOAc 8/1) to afford 2d (330 mg, 1.48 mmol, 98%) as a yellow solid. Mp: 109–110 °C. Rf (pentane/EtOAc 8/1): 0.3. \textsuperscript{1}H NMR (400 MHz, CDCl\textsubscript{3}) \(\delta 7.46 (dt, 2\ H, J = 9.5, 2.6\ Hz, \text{ArH}), 7.35 (ddd, 1\ H, J = 7.5, 1.6, 0.6\ Hz, \text{ArH}), 7.12 (ddd, 1\ H, J = 8.1, 7.3, 1.6\ Hz, \text{ArH}), 6.88 (dt, 2\ H, J = 9.4, 2.6\ Hz,
ArH), 6.71 (m, 2 H, ArH), 4.25 (m, 2 H, NH₂), 3.83 (s, 3 H, CH₃). ¹³C NMR (100 MHZ, CDCl₃) δ 159.6, 147.6, 132.9, 132.0, 129.4, 118.0, 115.4, 114.3, 114.0, 108.3, 94.6, 84.4, 55.3. Consistent with reported values [5].

4-Chloro-2-((4-fluorophenyl)ethynyl)aniline (2e)

Following a slightly modified procedure [5], a solution of 4-chloro-2-iodoaniline (4b) (304 mg, 1.20 mmol, 1 equiv), 1-ethynyl-4-fluorobenzene (12) (173 mg, 1.44 mmol, 1.2 equiv), PdCl₂(PPh₃)₂ (84 mg, 0.12 mmol, 0.1 equiv) and CuI (23 mg, 0.12 mmol, 0.1 equiv) was heated under reflux in Et₃N (12 mL) for 2 h under a nitrogen atmosphere. The resulting mixture was filtered through Celite®, washed with DCM and concentrated under vacuum. The resulting solid was purified by column chromatography (pentane/Et₂O 8/2 to 6/4) to afford 2e (283 mg, 1.15 mmol, 96%) as a yellow solid. Mp: 84–86 °C. Rf (pentane/Et₂O 8/2): 0.3. ¹H NMR (400 MHZ, CDCl₃) δ 7.50 (m, 2 H, ArH), 7.32 (d, 1 H, J = 2.4 Hz, ArH), 7.07 (m, 3 H, ArH), 6.65 (d, 1 H, J = 8.7 Hz, ArH), 4.26 (s, 2 H, NH₂). ¹³C NMR (100 MHZ, CDCl₃) δ 162.7 (d, J = 250 Hz), 146.4, 133.5 (d, J = 8 Hz), 131.4, 129.8, 122.3, 119.0, 115.9, 115.8 (d, J = 22 Hz), 109.1, 94.5, 84.4. IR ν 3474 (w), 3385 (w), 3050 (w), 1890 (w), 1612 (m), 1508 (s), 1486 (s), 1411 (m), 1309 (w), 1284 (w), 1229 (s), 1155 (m), 1092 (w), 901 (w), 834 (s), 810 (s), 787 (m), 741 (w), 680 (w). HRMS (ESI) calcd for C₁₄ClFH₁₀N⁺ [M+H]⁺ 246.0480; found 246.0484.

4-Chloro-2-((4-methoxyphenyl)ethynyl)aniline (2f)

Following a slightly modified procedure [5], a solution of 4-chloro-2-iodoaniline (4b) (304 mg, 1.20 mmol, 1 equiv), 1-ethynyl-4-methoxybenzene (13) (206 mg, 1.56 mmol, 1.3 equiv), PdCl₂(PPh₃)₂ (84 mg, 0.12 mmol, 0.1 equiv) and CuI (23 mg, 0.12 mmol, 0.1 equiv) were heated under reflux in Et₃N (12 mL) for 2 h under a nitrogen atmosphere. The resulting mixture was filtered through Celite®, washed with DCM and concentrated under vacuum. The resulting solid was purified by column chromatography (pentane/EtOAc 8/2) to afford 2f (331 mg, 1.33 mmol, 89%) as an orange solid. Mp: 97–99 °C. Rf (pentane/EtOAc
8/2): 0.3. \( ^1\)H NMR (400 MHz, CDCl\(_3\)) \( \delta \) 7.46 (m, 2 H, ArH), 7.32 (d, 1 H, \( J = 2.5 \) Hz, ArH), 7.07 (dd, 1 H, \( J = 8.6, 2.5 \) Hz, ArH), 6.89 (m, 2 H, ArH), 6.64 (d, 1 H, \( J = 8.7 \) Hz, ArH), 4.26 (s, 2 H, NH\(_2\)), 3.83 (s, 3 H, CH\(_3\)). \( ^{13}\)C NMR (100 MHz, CDCl\(_3\)) \( \delta \) 159.8, 146.2, 133.0, 131.2, 129.3, 122.2, 115.3, 114.8, 114.1, 109.7, 95.6, 83.3, 55.3. IR \( \nu \) 3464 (w), 3362 (w), 3037 (w), 2932 (w), 2840 (w), 2201 (w), 1605 (s), 1567 (w), 1513 (s), 1488 (s), 1460 (w), 1410 (w), 1294 (s), 1249 (s), 1174 (m), 1152 (w), 1089 (w), 1032 (m), 901 (w), 839 (s), 818 (m), 778 (w), 737 (m).

HRMS (ESI) calcd for C\(_{15}\)H\(_{13}\)ClN\(_2\)O\(_2\)\([M+H]^+\) 258.0686; found 258.0683.

**4-Amino-3-((4-methoxyphenyl)ethynyl)benzonitrile (2g)**

Following a slightly modified procedure [5], a solution of 4-cyano-2-iodoaniline (4c) (366 mg, 1.50 mmol, 1 equiv), 1-ethynyl-4-methoxybenzene (13) (238 mg, 1.80 mmol, 1.2 equiv), PdCl\(_2\)(PPh\(_3\))\(_2\) (102 mg, 0.15 mmol, 0.1 equiv) and CuI (28 mg, 0.15 mmol, 0.1 equiv) were heated under reflux in Et\(_3\)N (15 mL) for 2 h under a nitrogen atmosphere. The resulting mixture was filtered through Celite\(^{\circledR}\), washed with DCM and concentrated under vacuum. The resulting solid was purified by column chromatography (pentane/EtOAc 7/3) to afford 2g (331 mg, 1.33 mmol, 89%) as an orange solid. Mp: 138–140 °C. Rf (pentane/EtOAc 7/3): 0.3. \( ^1\)H NMR (400 MHz, CDCl\(_3\)) \( \delta \) 7.60 (d, 1 H, \( J = 1.5 \) Hz, ArH), 7.46 (d, 2 H, \( J = 8.7 \) Hz, ArH), 7.34 (dd, 1 H, \( J = 8.5, 1.6 \) Hz, ArH), 6.89 (d, 2 H, \( J = 8.7 \) Hz, ArH), 6.70 (d, 1 H, \( J = 8.5 \) Hz, ArH), 4.79 (s, 2 H, NH\(_2\)), 3.83 (s, 3 H, CH\(_3\)). \( ^{13}\)C NMR (100 MHz, CDCl\(_3\)) \( \delta \) 160.0, 150.8, 136.1, 133.0, 132.9, 119.4, 114.3, 114.1, 113.8, 108.5, 100.0, 96.2, 82.0, 55.3. IR \( \nu \) 3459 (m), 3354 (m), 3218 (w), 3055 (m), 2956 (w), 2838 (w), 2217 (s), 1618 (s), 1606 (s), 1559 (w), 1505 (s), 1467 (w), 1423 (w), 1337 (w), 1288 (m), 1248 (s), 1174 (m), 1158 (w), 1109 (w), 1026 (m), 907 (w), 832 (s), 787 (w), 736 (w). HRMS (ESI) calcd for C\(_{16}\)H\(_{13}\)N\(_2\)O\(_2\)\([M+H]^+\) 249.1022; found 249.1014.

**2-(Hex-1-yn-1-yl)aniline (2h)**

Following a slightly modified procedure [5], a solution of 2-iodoaniline (4a) (438 mg, 2.00 mmol, 1 equiv), 1-hexyne (14) (277 μL, 2.4 mmol, 1.2 equiv), PdCl\(_2\)(PPh\(_3\))\(_2\) (135 mg, 0.200 mmol, 0.1 equiv) and CuI (38 mg, 0.20 mmol, 0.1 equiv) were heated under reflux in
Et₃N (20 mL) for 2 h under a nitrogen atmosphere. The resulting mixture was filtered through Celite®, washed with DCM and concentrated under vacuum. The resulting solid was purified by column chromatography (pentane/EtOAc 98/2) to afford 2h (293 mg, 1.69 mmol, 85%) as a yellow liquid. Rf (pentane/EtOAc 98/2): 0.15. ¹H NMR (400 MHz, CDCl₃) δ 7.26 (dd, 1 H, J = 7.5, 1.2 Hz, ArH), 7.09 (m, 1 H, ArH), 6.68 (m, 2 H, ArH), 4.17 (br s, 2 H, NH₂), 2.49 (t, 2 H, J = 7.0 Hz, CH₂), 1.62 (m, 2 H, CH₂), 1.51 (m, 2 H, CH₂), 0.97 (t, 3 H, J = 7.3 Hz, CH₃). ¹³C NMR (100 MHz, CDCl₃) δ 147.7, 132.1, 128.8, 117.9, 114.2, 109.0, 95.8, 77.0, 31.1, 22.1, 19.4, 13.7. Consistent with reported values [6].

2-((Trimethylsilyl)ethynyl)aniline (2i)

Following a slightly modified procedure [5], a solution of 2-iodoaniline (4a) (438 mg, 2.00 mmol, 1 equiv), trimethylsilyl acetylene (15) (342 μL, 2.40 mmol, 1.2 equiv), PdCl₂(PPh₃)₂ (135 mg, 0.200 mmol, 0.1 equiv) and CuI (38 mg, 0.20 mmol, 0.1 equiv) were heated under reflux in Et₃N (20 mL) for 2 h under a nitrogen atmosphere. The resulting mixture was filtered through Celite®, washed with DCM and concentrated under vacuum. The resulting solid was purified by column chromatography (pentane/EtOAc 98/2) to afford 2i (242 mg, 1.28 mmol, 64%) as a colorless liquid. Rf (pentane/EtOAc 98/2): 0.25. ¹H NMR (400 MHz, CDCl₃) δ 7.31 (dd, J = 7.5, 1.2 Hz, 1 H, ArH), 7.12 (dt, J = 7.8, 1.6 Hz, ArH), 6.67 (m, 2 H, ArH), 4.24 (br s, 2 H, NH₂), 0.29 (s, 9 H, TMS). ¹³C NMR (100 MHz, CDCl₃) δ 148.2, 132.2, 129.8, 117.7, 114.1, 107.7, 101.8, 99.7, 0.1. Consistent with reported values [7].

2-(Ethynyl)aniline (2j)

KOH (1M in H₂O, 700 μL, 0.700 mmol, 1.1 equiv) was added to a stirred solution of 2i (121 mg, 0.640 mmol, 1 equiv) in MeOH (2 mL). After 1 h, the reaction was diluted in DCM (20 mL) and water (20 mL). The layers were separated and the aqueous layer extracted with DCM (20 mL). The organic layers were combined, washed with brine (20 mL), dried over MgSO₄, filtered and concentrated under vacuum to afford 2j (69 mg, 0.59 mmol, 92%) as a yellow oil. ¹H NMR (400 MHz, CDCl₃) δ 7.35 (dd, 1 H, J = 8.0, 1.6 Hz, ArH), 7.16 (td, J = 7.8, 1.5 Hz, 1 H, ArH), 6.70 (m, 2 H, ArH), 4.26 (br s, 2 H, NH₂), 3.40 (s, 1 H, CH).
Sequential annulations/direct alkynylation

2-Phenyl-3-((triisopropylsilyl)ethynyl)-1H-indole (3a)

NaAuCl₄ (3.2 mg, 0.0081 mmol, 0.02 equiv) was added to a stirred solution of 2a (78 mg, 0.40 mmol, 1 equiv) in iPrOH (3 mL) under an ambient atmosphere. The reaction was stirred at RT for 3 h. TIPS-EBX (1) (206 mg, 0.480 mmol, 1.2 equiv) and then AuCl (3.7 mg, 0.016 mmol, 0.04 equiv) were added. The reaction was stirred for 30 h and then concentrated under vacuum. Et₂O (20 mL) was added, the organic layer washed twice with 0.1 M NaOH (20 mL). The aqueous layers were combined and extracted with Et₂O (20 mL). The organic layers were combined, washed successively with saturated NaHCO₃ (20 mL) and brine (20 mL), dried with MgSO₄ and concentrated under reduced pressure. Purification by flash chromatography (pentane/Et₂O 8/2) afforded 3a (1. run: 145 mg, 0.388 mmol, 97%, 2. run: 142 mg, 0.380 mmol, 95%) as a brown amorphous solid. Rf (pentane/Et₂O 8/2): 0.4. ¹H NMR (CDCl₃, 400 MHz) δ 8.32 (br s, 1 H; NH), 8.11 (m, 2 H; ArH), 7.80 (m, 1 H; ArH), 7.49 (m, 2 H; ArH), 7.40 (m, 2 H; ArH), 7.40 (m, 2 H; ArH), 7.27 (m, 2 H; ArH), 1.24 (m, 21 H; TIPS). ¹³C NMR (CDCl₃, 100 MHz) δ 139.7, 135.1, 131.3, 130.8, 128.7, 128.3, 126.4, 123.4, 120.9, 120.1, 110.9, 101.2, 96.5, 95.1, 18.8, 11.5. Consistent with reported values [9].

2-(p-Tolyl)-3-((triisopropylsilyl)ethynyl)-1H-indole (3b)

NaAuCl₄ (3.2 mg, 0.0081 mmol, 0.02 equiv) was added to a stirred solution of 2b (83 mg, 0.40 mmol, 1 equiv) in iPrOH (3 mL) under an ambient atmosphere. The reaction was stirred at RT for 3 h. TIPS-EBX (1) (206 mg, 0.480 mmol, 1.2 equiv) and then AuCl (3.7 mg, 0.016 mmol, 0.04 equiv) were added. The reaction was stirred for 24 h and then concentrated under vacuum. Et₂O (20 mL) was added, the organic layer washed twice with 0.1 M NaOH (20 mL)
mL). The aqueous layers were combined and extracted with Et₂O (20 mL). The organic layers were combined, washed successively with saturated NaHCO₃ (20 mL) and brine (20 mL), dried with MgSO₄ and concentrated under reduced pressure. Purification by flash chromatography (pentane/Et₂O 9/1) afforded 3b (115 mg, 0.296 mmol, 74%) as an orange amorphous solid. Rf (pentane/Et₂O 9/1): 0.2. ¹H NMR (400 MHz, CDCl₃) δ 8.26 (s, 1 H, NH), 8.00 (d, 2 H, J = 8.2 Hz, ArH), 7.75 (d, 1 H, J = 6.9 Hz, ArH), 7.36 (m, 1 H, ArH), 7.25 (m, 4 H, ArH), 2.42 (s, 3 H, CH₃), 1.21 (m, 21 H, TIPS).

¹³C NMR (100 MHz, CDCl₃) δ 139.8, 138.4, 135.0, 130.9, 129.4, 128.6, 126.2, 123.3, 120.9, 120.0, 110.8, 101.4, 96.0, 95.0, 21.4, 18.8, 11.5. IR ν 3418 (m), 3063 (w), 2942 (s), 2863 (s), 2141 (s), 1676 (w), 1617 (w), 1504 (w), 1458 (s), 1382 (w), 1327 (m), 1305 (w), 1235 (m), 1175 (w), 1115 (w), 1060 (w), 1010 (w), 997 (m), 910 (m), 883 (m), 821 (m), 780 (m), 743 (s), 677 (s). HRMS (ESI) calcd for C₂₆H₃₄NSi⁺ [M+H]⁺ 388.2455; found 388.2459

2-(4-Fluorophenyl)-3-((triisopropylsilyl)ethynyl)-1H-indole (3c)

NaAuCl₄ (3.2 mg, 0.0081 mmol, 0.02 equiv) was added to a stirred solution of 2c (84 mg, 0.40 mmol, 1 equiv) in iPrOH (3 mL) under an ambient atmosphere. The reaction was stirred at RT for 3 h. TIPS-EBX (I) (206 mg, 0.480 mmol, 1.2 equiv) and then AuCl (3.7 mg, 0.016 mmol, 0.04 equiv) were added. The reaction was stirred for 30 h and then concentrated under vacuum. Et₂O (20 mL) was added, the organic layer washed twice with 0.1 M NaOH (20 mL). The aqueous layers were combined and extracted with Et₂O (20 mL). The organic layers were combined, washed successively with saturated NaHCO₃ (20 mL) and brine (20 mL), dried with MgSO₄ and concentrated under reduced pressure. Purification by flash chromatography (pentane/Et₂O 8/2) afforded 3c (124 mg, 0.317 mmol, 79%) as brown amorphous solid. Rf (pentane/Et₂O 8/2): 0.4. ¹H NMR (400 MHz, CDCl₃) δ 8.08 (s, 1 H, NH), 7.93 (dd, 2 H, J = 8.9, 5.3 Hz, ArH), 7.65 (m, 1 H, ArH), 7.25 (m, 1 H, ArH), 7.14 (m, 2 H, ArH), 7.04 (t, 2 H, J = 8.6 Hz, ArH), 1.10 (m, 21 H, TIPS). ¹³C NMR (100 MHZ, CDCl₃) δ 162.7 (d, J = 249 Hz), 138.8, 135.1, 130.7, 128.4 (d, J = 8 Hz), 127.7 (d, J = 3 Hz), 123.6, 121.1, 120.1, 115.9 (d, J = 22 Hz), 110.9, 101.0, 96.5, 95.2, 18.8, 11.5. IR ν 3422 (w), 2864 (m), 1890 (w), 1546 (w), 1502 (m), 1457 (m), 1440 (m), 1367 (w), 1327 (m), 1235 (s), 1162 (m), 1153 (w), 1104 (w), 1059 (w), 996 (m), 908 (m), 883 (m), 821 (s), 781 (m), 781 (m), 742 (s), 679 (s). HRMS (ESI) calcd for C₂₅FH₃₁NSi⁺ [M+H]⁺ 392.2204; found 392.2195
2-(4-Methoxyphenyl)-3-((triisopropylsilyl)ethynyl)-1H-indole (3d)

NaAuCl₄ (3.2 mg, 0.0081 mmol, 0.02 equiv) was added to a stirred solution of 2d (89 mg, 0.40 mmol, 1 equiv) in iPrOH (3 mL) under an ambient atmosphere. The reaction was stirred at RT for 3 h. DCM (1.5 mL), TIPS-EBX (1) (206 mg, 0.480 mmol, 1.2 equiv) and then AuCl (3.7 mg, 0.016 mmol, 0.04 equiv) were added. The reaction was stirred for 18 h and then concentrated under vacuum. Et₂O (20 mL) was added, the organic layer washed twice with 0.1 M NaOH (20 mL). The aqueous layers were combined and extracted with Et₂O (20 mL). The organic layers were combined, washed successively with saturated NaHCO₃ (20 mL) and brine (20 mL), dried with MgSO₄ and concentrated under reduced pressure. Purification by flash chromatography (pentane/Et₂O 8/2) afforded 3d (128 mg, 0.317 mmol, 79%) as a brown oil. Rf (pentane/Et₂O 8/2): 0.4. ¹H NMR (400 MHz, CDCl₃) δ 8.22 (s, 1 H, NH), 8.04 (m, 2 H, ArH), 7.75 (m, 1 H, ArH), 7.36 (m, 1 H, ArH), 7.24 (m, 2 H, ArH), 7.00 (m, 2 H, ArH), 3.90 (s, 3 H, CH₃), 1.22 (m, 21H, TIPS). ¹³C NMR (100 MHz, CDCl₃) δ 159.7, 139.9, 135.0, 130.9, 129.6, 127.8, 124.1, 123.1, 119.9, 114.1, 110.7, 101.5, 95.4, 94.7, 55.4, 18.8, 11.5. IR ν 3415 (w), 3063 (w), 2863 (m), 2140 (m), 1612 (m), 1578 (w), 1545 (w), 1504 (s), 1458 (s), 1439 (m), 1367 (w), 1328 (w), 1308 (m), 1285 (m), 1255 (s), 1184 (m), 1116 (w), 1062 (w), 1032 (m), 1019 (w), 911 (w), 833 (m), 834 (m), 790 (m), 744 (s), 678 (m), 659 (m). HRMS (ESI) calcd for C₂₆H₃₄NOSi⁺ [M+H]⁺ 404.2404; found 404.2423.

5-Chloro-2-(4-fluorophenyl)-3-((triisopropylsilyl)ethynyl)-1H-indole (3e)

NaAuCl₄ (3.2 mg, 0.0081 mmol, 0.02 equiv) was added to a stirred solution of 2e (98 mg, 0.40 mmol, 1 equiv) in iPrOH (3 mL) under an ambient atmosphere. The reaction was stirred at RT for 3 h. TIPS-EBX (1) (206 mg, 0.480 mmol, 1.2 equiv) and then AuCl (3.7 mg, 0.016 mmol, 0.04 equiv) were added. The reaction was stirred for 30 h and then concentrated under vacuum. Et₂O (20 mL) was added, the organic layer washed twice with 0.1 M NaOH (20 mL). The aqueous layers were combined and extracted with Et₂O (20 mL). The organic layers were combined, washed successively with saturated NaHCO₃ (20 mL) and brine (20 mL),
dried with MgSO₄ and concentrated under reduced pressure. Purification by flash chromatography (pentane/Et₂O 9/1 to 7/3) afforded 3e (124 mg, 0.291 mmol, 73%) as a grey solid. Mp: 99–100 °C. Rf (pentane/Et₂O 9/1): 0.2. ¹H NMR (400 MHz, CDCl₃) δ 8.25 (s, 1 H, NH), 8.03 (m, 2 H, ArH), 7.68 (d, 1 H, J = 1.5 Hz, ArH), 7.27 (m, 1 H, ArH), 7.18 (m, 3 H, ArH), 1.21 (m, 21 H, TIPS). ¹³C NMR (100 MHz, CDCl₃) δ 162.8 (d, J = 249 Hz), 140.1, 133.4, 131.7, 128.4 (d, J = 8 Hz), 127.2 (d, J = 3 Hz), 126.9, 123.9, 119.6, 115.9 (d, J = 22 Hz), 112.0, 100.1, 96.2, 95.8, 18.8, 11.4. IR ν 2926 (s), 2848 (s), 2106 (w), 1587 (w), 1530 (w), 1486 (m), 1433 (s), 1344 (w), 1294 (w), 1260 (m), 1200 (w), 1141 (w), 1089 (w), 1052 (w), 961 (w), 893 (s), 863 (m), 827 (m), 774 (w), 759 (m), 707 (s). HRMS (ESI) calcd for C₂₅ClF₃₀NSi⁺ [M+H]+ 426.1815; found 426.1824.

5-Chloro-2-(4-methoxyphenyl)-3-((triisopropylsilyl)ethynyl)-1H-indole (3f)

NaAuCl₄ (3.2 mg, 0.0081 mmol, 0.02 equiv) was added to a stirred solution of 2f (103 mg, 0.400 mmol, 1 equiv) in iPrOH (3 mL) under an ambient atmosphere. The reaction was stirred at RT for 3 h. DCM (1.5 mL), TIPS-EBX (1) (206 mg, 0.480 mmol, 1.2 equiv) and then AuCl (7.4 mg, 0.032 mmol, 0.08 equiv) were added. The reaction was stirred for 30 h and then concentrated under vacuum. Et₂O (20 mL) was added, the organic layer washed twice with 0.1 M NaOH (20 mL). The aqueous layers were combined and extracted with Et₂O (20 mL). The organic layers were combined, washed successively with saturated NaHCO₃ (20 mL) and brine (20 mL), dried with MgSO₄ and concentrated under reduced pressure. Purification by flash chromatography (pentane/Et₂O 8/2) afforded 3f (99 mg, 0.23 mmol, 56%, 90% pure) as a grey solid. Analytically pure product was obtained by preparative TLC (pentane/Et₂O 7/3). Rf (pentane/Et₂O 7/3): 0.2. Mp: 180–182 °C. ¹H NMR (400 MHz, CDCl₃) δ 8.24 (s, 1 H, NH), 8.01 (m, 2 H, ArH), 7.65 (d, 1 H, J = 2.0 Hz, ArH), 7.27 (m, 1 H, ArH), 7.18 (dd, 1 H, J = 8.5, 2.0 Hz, ArH), 6.99 (m, 2 H, ArH), 3.88 (s, 3 H, CH₃), 1.19 (m, 21 H, TIPS). ¹³C NMR (100 MHz, CDCl₃) δ 160.0, 141.2, 133.3, 132.0, 127.9, 126.7, 123.6, 123.4, 119.4, 114.2, 111.7, 100.6, 98.4, 95.3, 55.4, 18.8, 11.5. IR ν 3427 (w), 2865 (w), 2141 (w), 1545 (w), 1503 (m), 1467 (s), 1311 (w), 1288 (m), 1256 (s), 1183 (m), 1073 (w), 1033 (m), 997 (w), 910 (m), 883 (w), 837 (m), 803 (s), 735 (m), 697 (m), 667 (m). HRMS (ESI) calcd for C₂₆ClH₃₃NOSi⁺ [M+H]+ 438.2014; found 438.2018.
2-(4-Methoxyphenyl)-3-((triisopropylsilyl)ethynyl)-1H-indole-5-carbonitrile (3g)

NaAuCl₄ (6.4 mg, 0.016 mmol, 0.04 equiv) was added to a stirred solution of 2g (99 mg, 0.40 mmol, 1 equiv) in iPrOH (3 mL) under an ambient atmosphere. The reaction was stirred at 80°C for 3 h. The reaction was cooled to RT and DCM (1.5 mL), TIPS-EBX (1) (206 mg, 0.480 mmol, 1.2 equiv) and then AuCl (7.4 mg, 0.032 mmol, 0.08 equiv) were added. The reaction was stirred for 18 h and then TIPS-EBX (206 mg, 0.480 mmol, 1.2 equiv) added. After 12 h the reaction was concentrated under vacuum. Et₂O (20 mL) was added, the organic layer washed twice with 0.1 M NaOH (20 mL). The aqueous layers were combined and extracted with Et₂O (20 mL). The organic layers were combined, washed successively with saturated NaHCO₃ (20 mL) and brine (20 mL), dried with MgSO₄ and concentrated under reduced pressure. Purification by flash chromatography (pentane/Et₂O 5/5 to 3/7) afforded 3g (98 mg, 0.23 mmol, 54%, 95% pure) as a grey solid. A second batch of product 3g (48 mg, 0.11 mmol, 25%, 90% pure) was also obtained. Analytically pure product was obtained by recrystallization from hexanes/EtOAc. Combined yield: 79%. Mp: 158–159 °C. ¹H NMR (400 MHz, CDCl₃) δ 8.97 (s, 1 H, NH), 8.04 (m, 2 H, ArH), 8.00 (m, 1 H, ArH), 7.42 (m, 2 H, ArH), 6.97 (m, 2 H, ArH), 3.85 (m, 3 H, CH₃), 1.20 (m, 21 H, TIPS). ¹³C NMR (100 MHZ, CDCl₃) δ 160.3, 142.0, 136.7, 130.7, 128.1, 126.0, 125.0, 123.0, 120.7, 114.3, 111.7, 103.6, 99.8, 96.4, 95.7, 55.4, 18.8, 11.4. IR ν 3310 (m), 2941 (m), 2864 (m), 2223 (m), 1707 (w), 1615 (m), 1586 (w), 1546 (w), 1504 (s), 1473 (s), 1382 (w), 1293 (m), 1255 (s), 1184 (s), 1133 (w), 1034 (m), 1025 (w), 909 (m), 884 (m), 833 (m), 805 (m), 739 (s), 672 (m), 661 (m). HRMS (ESI) calcd for C₂₇H₃₃N₂OSi⁺ [M+H]⁺ 429.2357; found 429.2369.

2-Butyl-3-((triisopropylsilyl)ethynyl)-1H-indole (3h)

NaAuCl₄ (3.2 mg, 0.0081 mmol, 0.02 equiv) was added to a stirred solution of 2h (69 mg, 0.40 mmol, 1 equiv) in iPrOH (3 mL) under an ambient atmosphere. The reaction was stirred at RT for 3 h. TIPS-EBX (1) (206 mg, 0.480 mmol, 1.2 equiv) and then AuCl (3.7 mg, 0.016
mmol, 0.04 equiv) were added. The reaction was stirred for 4 h and the concentrated under vacuum. Et$_2$O (20 mL) was added, the organic layer washed twice with 0.1 M NaOH (20 mL). The aqueous layers were combined and extracted with Et$_2$O (20 mL). The organic layers were combined, washed successively with saturated NaHCO$_3$ (20 mL) and brine (20 mL), dried with MgSO$_4$ and concentrated under reduced pressure. Purification by flash chromatography (pentane/Et$_2$O 9/1) afforded 3h (120 mg, 0.339 mmol, 85%) as yellow oil. Rf (pentane/Et$_2$O 9/1): 0.4. $^1$H NMR (400 MHz, CDCl$_3$) δ 7.88 (s, 1 H, NH), 7.65 (m, 1 H, ArH), 7.27 (m, 1H, ArH), 7.16 (m, 2 H, ArH), 2.90 (t, 2 H, $J = 7.5$ Hz, CH$_2$), 1.74 (m, 2 H, CH$_2$), 1.41 (m, 2 H, CH$_2$), 1.18 (m, 21 H, TIPS), 0.95 (t, 3 H, $J = 7.3$ Hz, CH$_3$). $^{13}$C NMR (100 MHZ, CDCl$_3$) δ 144.7, 134.4, 129.5, 122.1, 120.5, 119.4, 110.5, 100.6, 96.7, 93.1, 31.2, 27.1, 22.3, 18.8, 13.8, 11.4. IR ν 3401 (m), 2958 (s), 2942 (s), 2865 (s), 2146 (s), 1617 (w), 1548 (w), 1460 (s), 1382 (w), 1330 (w), 1242 (m), 1158 (w), 1077 (w), 1000 (w), 997 (w), 920 (w), 883 (m), 778 (m), 743 (s), 677 (s), 631 (s). HRMS (ESI) calcd for C$_{23}$H$_{36}$NSi$^+$ [M+H]$^+$ 354.2612; found 354.2606.

References

Spectra of new compounds

solvent: CDCl₃
Frequency: 400.13 MHz

solvent: CDCl₃
Frequency: 100.613 MHz

solvent: CDCl₃
Frequency: 100.613 MHz
solvent: CDCl3
Frequency: 400.13 MHz

solvent: CDCl3
Frequency: 100.61 MHz

solvent: CDCl3
Frequency: 100 MHz

solvent: CDCl3
Frequency: 100 MHz