


Cite this: *RSC Adv.*, 2024, 14, 1106

Received 8th November 2023

Accepted 4th December 2023

DOI: 10.1039/d3ra07636a

rsc.li/rsc-advances

Enantioselective synthesis of α -tetrasubstituted (1-indoliziny) (diaryl)-methanamines *via* chiral phosphoric acid catalysis†

Jialing Zhong, Rihuang Pan and Xufeng Lin *

An enantioselective Friedel–Crafts reaction of cyclic α -diaryl *N*-acyl imines with indolizines catalyzed by a chiral spirocyclic phosphoric acid has been developed. The asymmetric transformation proceeds smoothly to afford α -tetrasubstituted (1-indoliziny) (diaryl)methanamines in good yields with up to 98% ee under mild conditions.

Chiral α -tetrasubstituted methanamines are frequently distributed in diverse bioactive natural products,¹ and extensive effort has been devoted to constructing these scaffolds in synthetic chemistry over the past decade.² Some excellent chiral organocatalysts³ and chiral metal salt catalysts⁴ have been developed in the asymmetric synthesis of chiral α -tetrasubstituted (diaryl) alkyl or (triaryl) methanamines. Despite these notable advances, to the best of our knowledge, there are currently no versatile protocols for the asymmetric preparation of chiral α -(1-indoliziny)-(diaryl)methanamines.

Indolizines as an important class of N-containing heterocycles can be found in organic synthesis and numerous pharmaceuticals (Fig. 1).⁵ Many versatile strategies for the direct functionalization of indolizines have been developed.⁶ Moreover, some elegant examples toward the asymmetric synthesis of enantioenriched indolizine derivatives have been reported,⁷ as shown in Scheme 1. For instance, List and Coelho reported the first organocatalyzed asymmetric conjugate addition of indolizines to enones using the chiral BINOL-derived phosphoric acid (BINOL-PA) as a catalyst (Scheme 1a).^{7a} Zhang and Fu developed a copper-catalyzed enantioselective propargylation reaction of indolizines (Scheme 1b).^{7b} Later, Zeng's group established a highly asymmetric allylic substitution reaction of indolizine derivatives catalyzed by chiral Ir complexes (Scheme 1c).^{7c} Ni and Song described an organocatalytic highly diastereo- and enantioselective Friedel–Crafts conjugate addition of indolizines to prochiral cyclopentenones catalyzed by BINOL-PA (Scheme 1d).^{7d}

Recently, Li and Gu realized the catalytic asymmetric conjugate addition of indolizines to unsaturated ketones catalyzed by chiral Rh complexes (Scheme 1e).^{7e} Very recently, Ni

and Song reported BINOL-PA-catalyzed asymmetric atroposelective arylation of indolizines for the preparation of the axially chiral 3-arylindolizines (Scheme 1f).^{7f} Although these strategies enable direct access to asymmetric synthesis of chiral indolizine derivatives, development of new class indolizines is still a formidable target. To continue our efforts⁸ on the advancement of chiral phosphoric acid catalysis,⁹ we here present the chiral spirocyclic phosphoric acid (SPINOL-PA) catalyzed enantioselective Friedel–Crafts of indolizines with *in situ* generated cyclic α -diaryl *N*-acyl imines¹⁰ for the synthesis of chiral α -(1-indoliziny) (diaryl)methanamines.

An initial investigation for the asymmetric Friedel–Crafts reaction was carried out with methyl indolizine-2-carboxylate (**1a**) and 3-phenyl 3-hydroxyisindolinone (**2a**) in the presence of 10 mol% chiral phosphoric acid (CPA), as shown in Table 1. Chiral spirocyclic phosphoric acid (SPA) catalysts (**4a–d**) developed by our group^{8a} were firstly screened in 1,2-dichloroethane (DCE) at room temperature to afford the desired chiral α -(1-indoliziny) (diaryl)methanamine (**3a**) with acceptable yields but up to different enantioselectivities, and catalyst **4d** gave the best

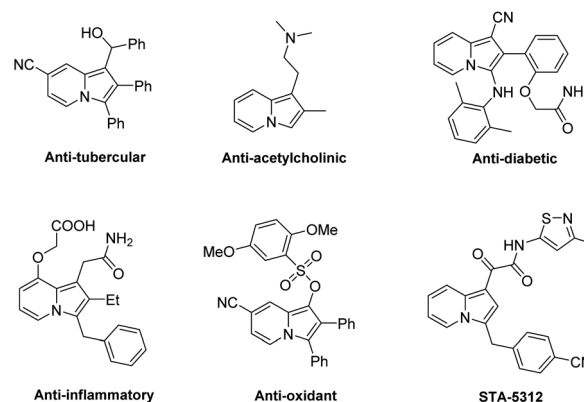


Fig. 1 Indolizine-containing bioactive compounds.

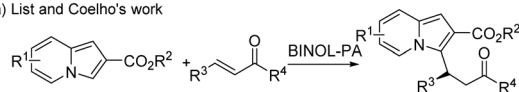
Department of Chemistry, Zhejiang University, Hangzhou 310027, P. R. China. E-mail: lxfoke@zju.edu.cn

† Electronic supplementary information (ESI) available. CCDC 2167990 (**3p**). For ESI and crystallographic data in CIF or other electronic format see DOI: <https://doi.org/10.1039/d3ra07636a>

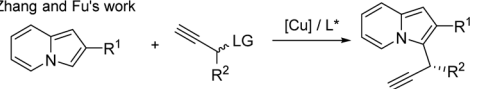


Previous work:

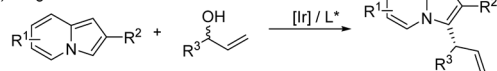
a) List and Coelho's work



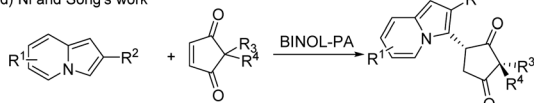
b) Zhang and Fu's work



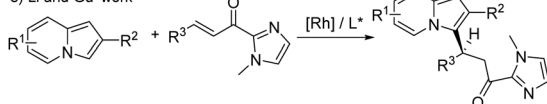
c) Zeng's work



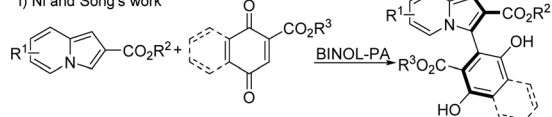
d) Ni and Song's work



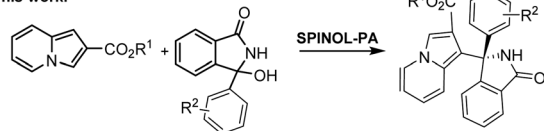
e) Li and Gu's work



f) Ni and Song's work



This work:



Scheme 1 Asymmetric functionalization of indolizines.

reaction activity and enantioselectivity (89% yield, 74% ee) (entries 1–4). In addition, we also tested BINOL-derived phosphoric acids (**5a–d**) as catalysts, only low yields (37–72%) and poor enantioselectivities (12–53% ee) were observed (entries 5–8). We believe that the efficient catalytic activity of chiral spirocyclic phosphoric acid (SPA) is due to its special skeleton. Mostly the chiral backbone of the catalyst plays a vital role in attaining high stereoselection by regulating electronic and structural properties of substrates. On the other hand, the dual hydrogen bonding network between the SPA and other two substrates plays a crucial role in terms of reactivity and selectivity. Moreover, the influence of solvent was investigated (entries 9–13). 1,2-Dichloroethane (DCE) was still the optimal solvent, and this reaction did not even proceed in toluene, THF or EtOAc. Next, the effect of additives was also investigated (entries 14–17). In the presence of 4 Å MS, the desired product **3a** could be obtained in 91% yield with 87% ee (17). Lastly, we examined the temperature, and the reaction proceeded for 36 hours to afford the product **3a** with 92% ee but in only 9% yield when the temperature was lowered to 0 °C (entry 18). Hence, the optimized reaction conditions employed 10 mol% (*S*)-**4d** in 1,2-dichloroethane at room temperature (entry 17).

With the optimal conditions in hand, we set out to explore the substrate scope and limitations of this asymmetric

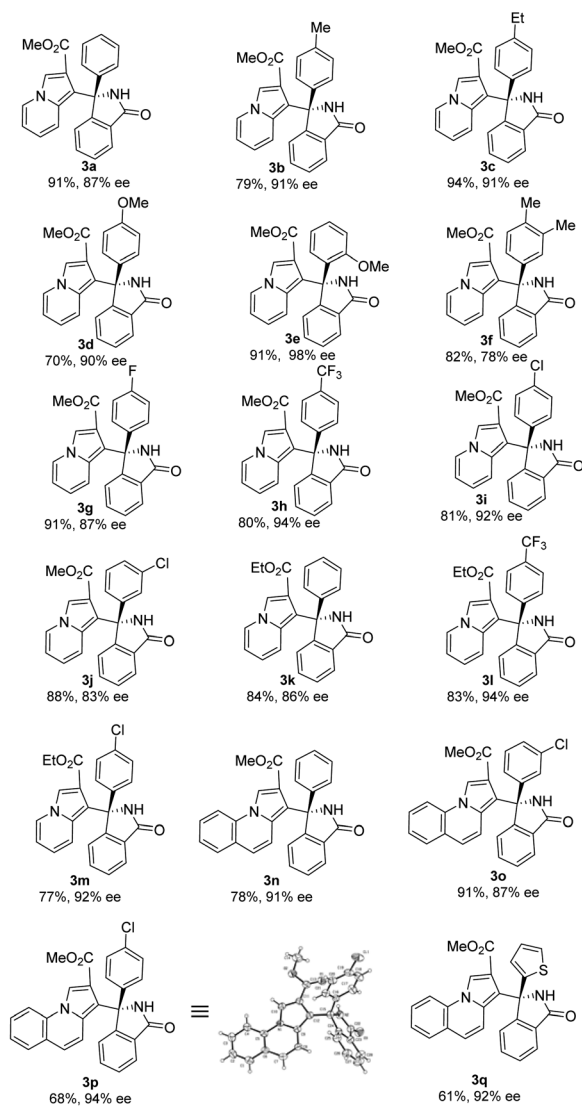
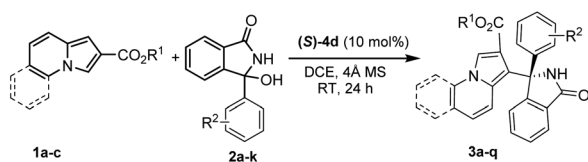
Table 1 Optimization of the reaction conditions^a

Entry	CPA	Solvent	Additive	Yield ^b (%)	ee ^c (%)
1	(<i>S</i>)- 4a	DCE	None	49	15
2	(<i>S</i>)- 4b	DCE	None	71	41
3	(<i>S</i>)- 4c	DCE	None	78	6
4	(<i>S</i>)- 4d	DCE	None	89	74
5	(<i>R</i>)- 5a	DCE	None	55	–12
6	(<i>R</i>)- 5b	DCE	None	37	–19
7	(<i>R</i>)- 5c	DCE	None	61	–23
8	(<i>R</i>)- 5d	DCE	None	72	–53
9	(<i>S</i>)- 4d	DCM	None	91	37
10	(<i>S</i>)- 4d	MeCN	None	55	23
11	(<i>S</i>)- 4d	Toluene	None	N.R.	—
12	(<i>S</i>)- 4d	THF	None	N.R.	—
13	(<i>S</i>)- 4d	EtOAc	None	N.R.	—
14	(<i>S</i>)- 4d	MeOH	None	N.R.	—
15	(<i>S</i>)- 4d	DCE	Na ₂ SO ₄	82	78
16	(<i>S</i>)- 4d	DCE	MgSO ₄	79	74
17	(<i>S</i>)- 4d	DCE	3 Å MS	80	84
18	(<i>S</i>)- 4d	DCE	4 Å MS	91	87
19 ^d	(<i>S</i>)- 4d	DCE	4 Å MS	9	91

^a Reactions were performed with **1a** (0.05 mmol), **2a** (0.05 mmol) and CPA catalyst (10 mol%) in the presence of additive (100 mg) in solvent (1 mL) for 24 hours at room temperature. ^b Isolated yields. ^c Determined by chiral HPLC analysis. ^d At 0 °C for 36 hours.

transformation, as summarized in Table 2. In general, a range of 3-aryl 3-hydroxyisoindolinones **2** with different substituents were amenable to this strategy, and reacted efficiently with methyl indolizine-2-carboxylate **1a** to provide good yields and high enantioselectivities (**3a–j**, up to 94% yield, up to 98% ee). When two methyl groups were placed around the 3-aryl ring, we observed a small drop in enantioselectivity as the corresponding product **3f** was obtained in 82% yield and 78% ee. Interestingly, when methoxy group was introduced in ortho position of the 3-aryl, the enantioselectivity was dramatically improved (**3e**, 91% yield, 98% ee). Furthermore, CF₃– group on the 3-aryl substituent proceeded smoothly to afford the desired product **3h** in good yield with excellent enantioselectivity (94% ee). We found that 3-alkyl 3-hydroxyisoindolinones were not tolerated and the reactions did not run with **1a** under the standard reaction conditions, such as methyl, allyl or benzyl substituent on the isoindolinone alcohol. We tried carrying out the

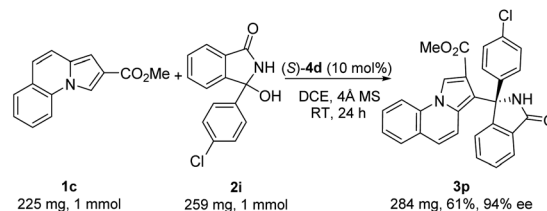


Table 2 Substrates scope^a

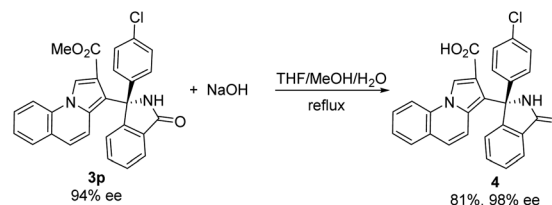
^a Reactions were performed with **1** (0.05 mmol), **2** (0.05 mmol) and (*S*)-**4d** (10 mol%) in the presence of 4 Å MS (100 mg) in DCE (1 mL) at room temperature for 24 hours. Isolated yield was given. The ee was determined by chiral HPLC analysis.

reactions of 3-alkyl 3-hydroxyisindolinones with substrate **1a** in HFIP, and found that the reactions still could not run.

We next examined the scope of indolizines **1**. Ethyl indolizine-2-carboxylate **1b** also gave the desired products (**3k–m**) in good yields with excellent enantioselectivities (up to 94% ee). However, indolizine with a 5-methyl or 5-phenyl substituent depressed the reactivity and failed to provide the desired



Scheme 2 1 mmol scale reaction.



Scheme 3 Derivatization experiment.

product under standard conditions because of the steric hindrance. Furthermore, methyl pyrrolo[1,2-*a*]quinoline-2-carboxylate **1c** also gave the desired products (**3n–q**) with high enantioselectivities (up to 94% ee). The absolute configuration in product **3p** (CCDC 2167990) was clearly determined to be (*R*) by X-ray diffraction analysis of a single crystal, and the absolute configuration of products **3** was assigned as (*R*) by analogy.

To further explore the synthetic practicality of the developed protocol, we investigated a 1 mmol scale reaction of this asymmetric Friedel–Crafts reaction, as shown in Scheme 2. Under the optimized reaction conditions, the reaction of methyl pyrrolo[1,2-*a*]quinoline-2-carboxylate **1c** (1 mmol) and 3-(4-chlorophenyl)-3-hydroxyisindolin-1-one **2i** (1 mmol) afforded the desired product **3p** in 61% yield with 94% ee.

We attempted to extend the reaction by treating product **3p** with NaOH and obtained the corresponding product **4** in 81% yield and 98% ee, as shown in Scheme 3.

Conclusions

In summary, we have reported a metal free protocol for chiral spirocyclic phosphoric acid-catalyzed enantioselective Friedel–Crafts reaction of cyclic α -diaryl *N*-acyl imines with indolizines, providing convenient access to a range of α -tetrasubstituted (1-indoliziny) (diaryl)methanamines in good yields with up to 98% ee under mild conditions.

Experimental

General information

All reactions were carried out in oven-dried glassware with magnetic stirring under ambient conditions. Unless otherwise noted, all reagents, including the chiral phosphoric acid catalysts **4** and **5**, were purchased from commercial supplies and used without further purification, and all solvents were dried and purified according to standard methods prior to use. Substrates **1** (ref. 11) and **2** (ref. 12) were synthesized according



to the literature methods. ^1H NMR and ^{13}C NMR spectra were recorded on a Bruker AVANCE III 400 MHz spectrometer instrument at 400 MHz and 100 MHz spectrometer, respectively. The chemical shifts (δ) were quoted in parts per million (ppm) downfield relative to internal standard TMS (0.0 ppm) and referenced to solvent peaks in the NMR solvent ($\text{CDCl}_3 = \delta$ 7.26 ppm; δ 77.00 ppm; $\text{D}_6\text{-DMSO} = \delta$ 2.50 ppm; δ 40.00 ppm). Spin multiplicity were reported using the following abbreviations: s = singlet, d = doublet, t = triplet, dd = doublet of doublet, td = triplet of doublet, m = multiplet. Infrared spectra were recorded on an ATR-FTIR spectrometer. ESI-HRMS were recorded on a Water Micromass GCT Premier mass spectrometer. Optical rotations were measured on a PerkinElmer Model 341 polarimeter at 20 °C. Enantiomeric excess (ee) were measured by chiral HPLC analysis.

General procedure for the asymmetric synthesis of **3** via chiral phosphoric acid-catalyzed reaction of indolizine-2-carboxylate **1** and quinone methyl ester **2**

To a mixture of indolizine-2-carboxylate **1** (0.05 mmol), 3-hydroxyisoindolinones **2** (0.05 mmol, 1 equiv.) and catalyst (*S*)-**4a** (0.005 mmol, 10 mol%) in DCE (1 mL) was added 4 Å MS (100 mg). After stirring at room temperature for 24 hours, the residue was purified by flash column chromatography with acetate/petroleum ether 1 : 2 (v/v) on silica gel to give the desired product **3**.

(R)-Methyl 1-(3-oxo-1-phenylisoindolin-1-yl) indolizine-2-carboxylate (3a). White solid (17.4 mg, 91%). Mp 119–120 °C. 87% ee, determined by HPLC [Daicel Chiralcel AD-H column (250 × 4.6 mm)], *n*-hexane/*i*-PrOH = 60/40, 1.0 mL min^{−1}, 254 nm, $t_{\text{major}} = 14.566$ min, $t_{\text{minor}} = 9.524$ min. $[\alpha]_{\text{D}}^{20} = -440.3^\circ$ (c 0.36, CH_2Cl_2). ^1H NMR (400 MHz, $\text{DMSO}-d_6$) δ 9.02 (s, 1H), 8.26 (d, $J = 6.9$ Hz, 1H), 8.08 (s, 1H), 7.76 (d, $J = 6.8$ Hz, 1H), 7.59–7.50 (m, 3H), 7.25 (m, 5H), 6.55 (m, 2H), 6.12 (d, $J = 9.3$ Hz, 1H), 3.25 (s, 3H) ppm. ^{13}C NMR (101 MHz, DMSO) δ 168.9, 165.3, 158.8, 151.1, 145.7, 132.3, 131.6, 130.4, 129.0, 128.6, 127.4, 127.2, 125.6, 125.5, 123.7, 119.6, 119.4, 119.2, 118.1, 112.3, 112.0, 67.1, 60.2, 51.5 ppm. IR (film): $\gamma = 3393$, 2949, 1694, 1610, 1541, 1503, 1466, 1370, 1313, 1266, 1222, 1159, 1075, 745, 702 cm^{−1}. HRMS m/z (ESI⁺): calcd for $\text{C}_{24}\text{H}_{18}\text{N}_2\text{NaO}_3$ ($[\text{M} + \text{Na}]^+$) 405.1210, found 405.1211.

(R)-Methyl 1-(3-oxo-1-(*p*-tolyl)isoindolin-1-yl)indolizine-2-carboxylate (3b). White solid (15.7 mg, 79%). Mp 118–120 °C. 91% ee, determined by HPLC [Daicel Chiralcel AD-H column (250 × 4.6 mm)], *n*-hexane/*i*-PrOH = 60/40, 1.0 mL min^{−1}, 254 nm, $t_{\text{major}} = 14.365$ min, $t_{\text{minor}} = 7.750$ min. $[\alpha]_{\text{D}}^{20} = -571.1^\circ$ (c 0.30, CH_2Cl_2). ^1H NMR (400 MHz, $\text{DMSO}-d_6$) δ 8.98 (s, 1H), 8.26 (d, $J = 6.8$ Hz, 1H), 8.07 (s, 1H), 7.75 (d, $J = 6.9$ Hz, 1H), 7.55–7.48 (m, 3H), 7.15 (d, $J = 8.2$ Hz, 2H), 7.06 (d, $J = 8.2$ Hz, 2H), 6.53 (m, 2H), 6.15 (d, $J = 9.3$ Hz, 1H), 3.27 (s, 3H), 2.23 (s, 3H) ppm. ^{13}C NMR (101 MHz, DMSO) δ 168.9, 165.4, 151.4, 142.7, 136.5, 132.3, 131.6, 130.4, 129.2, 128.9, 127.2, 125.6, 125.4, 123.7, 119.6, 119.4, 119.3, 118.0, 112.4, 112.0, 66.9, 51.6, 26.8, 21.2, 21.0 ppm. IR (film): $\gamma = 3403$, 2949, 1694, 1541, 1508, 1466, 1437, 1364, 1314, 1267, 1221, 1159, 1069, 815, 755, 692 cm^{−1}. HRMS m/z (ESI⁺): calcd for $\text{C}_{25}\text{H}_{20}\text{N}_2\text{NaO}_3$ ($[\text{M} + \text{Na}]^+$) 419.1366, found 419.1364.

(R)-Methyl 1-(1-(4-ethylphenyl)-3-oxoisoindolin-1-yl) indolizine-2-carboxylate (3c). White solid (19.3 mg, 94%). Mp 105–107 °C. 91% ee, determined by HPLC [Daicel Chiralcel AD-H column (250 × 4.6 mm)], *n*-hexane/*i*-PrOH = 60/40, 1.0 mL min^{−1}, 254 nm, $t_{\text{major}} = 14.463$ min, $t_{\text{minor}} = 7.273$ min. $[\alpha]_{\text{D}}^{20} = -486.6^\circ$ (c 0.38, CH_2Cl_2). ^1H NMR (400 MHz, $\text{DMSO}-d_6$) δ 8.99 (s, 1H), 8.26 (d, $J = 6.9$ Hz, 1H), 8.06 (s, 1H), 7.75 (d, $J = 6.8$ Hz, 1H), 7.55 (m, 2H), 7.53–7.48 (m, 1H), 7.17 (d, $J = 8.3$ Hz, 2H), 7.09 (d, $J = 8.3$ Hz, 2H), 6.59–6.47 (m, 2H), 6.15 (d, $J = 9.3$ Hz, 1H), 3.25 (s, 3H), 2.58–2.52 (m, 2H), 1.12 (t, $J = 7.6$ Hz, 3H) ppm. ^{13}C NMR (101 MHz, DMSO) δ 168.9, 165.4, 151.3, 142.9, 142.9, 132.3, 131.6, 130.3, 128.9, 128.0, 127.2, 125.7, 125.4, 123.7, 119.7, 119.4, 119.3, 117.9, 112.5, 111.9, 66.9, 51.6, 31.6, 30.3, 28.2, 16.1 ppm. IR (film): $\gamma = 3398$, 2961, 1694, 1610, 1540, 1504, 1466, 1437, 1364, 1313, 1267, 1221, 1159, 1097, 828, 753, 692 cm^{−1}. HRMS m/z (ESI⁺): calcd for $\text{C}_{26}\text{H}_{22}\text{N}_2\text{NaO}_3$ ($[\text{M} + \text{Na}]^+$) 433.1523, found 433.1526.

(R)-Methyl 1-(1-(4-methoxyphenyl)-3-oxoisoindolin-1-yl) indolizine-2-carboxylate (3d). White solid (14.4 mg, 70%). Mp 113–115 °C. 90% ee, determined by HPLC [Daicel Chiralcel AD-H column (250 × 4.6 mm)], *n*-hexane/*i*-PrOH = 60/40, 1.0 mL min^{−1}, 254 nm, $t_{\text{major}} = 12.837$ min, $t_{\text{minor}} = 6.504$ min. $[\alpha]_{\text{D}}^{20} = -434.1^\circ$ (c 0.26, CH_2Cl_2). ^1H NMR (400 MHz, $\text{DMSO}-d_6$) δ 9.01 (s, 1H), 8.25 (d, $J = 6.7$ Hz, 1H), 8.06 (s, 1H), 7.74 (d, $J = 7.1$ Hz, 1H), 7.56 (m, 2H), 7.53–7.48 (m, 1H), 7.18 (d, $J = 8.8$ Hz, 2H), 6.82 (d, $J = 8.8$ Hz, 2H), 6.54 (m, 2H), 6.19 (d, $J = 9.1$ Hz, 1H), 3.70 (s, 3H), 3.30 (s, 3H) ppm. ^{13}C NMR (101 MHz, DMSO) δ 168.8, 165.4, 158.6, 151.5, 137.6, 132.3, 131.6, 130.4, 128.8, 127.3, 127.2, 127.0, 125.3, 123.7, 119.6, 119.4, 117.9, 113.9, 112.6, 111.9, 100.0, 66.6, 60.3, 55.6, 51.7 ppm. IR (film): $\gamma = 3393$, 2950, 1694, 1608, 1508, 1466, 1437, 1370, 1313, 1297, 1266, 1221, 1177, 1096, 1033, 829, 745, 693 cm^{−1}. HRMS m/z (ESI⁺): calcd for $\text{C}_{25}\text{H}_{20}\text{N}_2\text{NaO}_4$ ($[\text{M} + \text{Na}]^+$) 435.1315, found 435.1317.

(R)-Methyl 1-(1-(2-methoxyphenyl)-3-oxoisoindolin-1-yl) indolizine-2-carboxylate (3e). White solid (18.8 mg, 91%). Mp 111–112 °C. 98% ee, determined by HPLC [Daicel Chiralcel AD-H column (250 × 4.6 mm)], *n*-hexane/*i*-PrOH = 60/40, 1.0 mL min^{−1}, 254 nm, $t_{\text{major}} = 8.571$ min, $t_{\text{minor}} = 9.692$ min. $[\alpha]_{\text{D}}^{20} = -412.3^\circ$ (c 0.39, CH_2Cl_2). ^1H NMR (500 MHz, CD_2Cl_2) δ 7.86–7.83 (m, 1H), 7.81 (d, $J = 6.7$ Hz, 1H), 7.73 (s, 1H), 7.64 (s, 1H), 7.61 (s, 1H), 7.53 (m, 2H), 7.28 (m, 3H), 6.94–6.84 (m, 2H), 6.43 (d, $J = 6.7$ Hz, 2H), 3.49 (s, 3H), 3.46 (s, 3H) ppm. ^{13}C NMR (101 MHz, DMSO) δ 169.4, 165.9, 157.4, 149.8, 131.65, 131.6, 129.4, 129.3, 128.6, 125.7, 125.5, 124.0, 120.3, 119.6, 119.0, 118.1, 116.1, 113.3, 112.6, 111.6, 65.7, 55.7, 51.8, 49.9, 29.7 ppm. IR (film): $\gamma = 3393$, 2949, 1694, 1610, 1541, 1503, 1466, 1370, 1313, 1266, 1222, 1159, 1075, 745, 702 cm^{−1}. HRMS m/z (ESI⁺): calcd for $\text{C}_{25}\text{H}_{20}\text{N}_2\text{NaO}_4$ ($[\text{M} + \text{Na}]^+$) 435.1315, found 435.1318.

(R)-Methyl 1-(1-(3,4-dimethylphenyl)-3-oxoisoindolin-1-yl) indolizine-2-carboxylate (3f). White solid (16.8 mg, 82%). Mp 113–114 °C. 78% ee, determined by HPLC [Daicel Chiralcel AD-H column (250 × 4.6 mm)], *n*-hexane/*i*-PrOH = 60/40, 1.0 mL min^{−1}, 254 nm, $t_{\text{major}} = 10.419$ min, $t_{\text{minor}} = 6.636$ min. $[\alpha]_{\text{D}}^{20} = -421.9^\circ$ (c 0.33, CH_2Cl_2). ^1H NMR (500 MHz, CDCl_3) δ 7.92 (d, $J = 6.6$ Hz, 1H), 7.79 (m, 3H), 7.54–7.43 (m, 2H), 7.34



(d, $J = 7.0$ Hz, 1H), 7.08 (s, 1H), 6.99 (m, 2H), 6.43 (t, $J = 6.7$ Hz, 1H), 6.40–6.30 (m, 1H), 5.88 (d, $J = 9.5$ Hz, 1H), 3.45 (s, 3H), 2.15 (d, $J = 9.1$ Hz, 6H) ppm. ^{13}C NMR (101 MHz, DMSO) δ 169.5, 165.6, 150.6, 142.2, 136.4, 135.3, 131.6, 131.0, 130.2, 129.5, 128.6, 126.0, 125.8, 125.7, 124.3, 122.5, 119.7, 119.6, 118.8, 117.6, 113.0, 111.8, 66.7, 51.6, 20.0, 19.4 ppm. IR (film): $\gamma = 3402, 2948, 1694, 1610, 1541, 1500, 1437, 1364, 1313, 1266, 1220, 1159, 1098, 816, 755, 692\text{ cm}^{-1}$. HRMS m/z (ESI^+): calcd for $\text{C}_{26}\text{H}_{22}\text{N}_2\text{NaO}_3$ ($[\text{M} + \text{Na}]^+$) 433.1523, found 433.1521.

(R)-Methyl 1-(1-(4-fluorophenyl)-3-oxoisindolin-1-yl)indolizine-2-carboxylate (3g). White solid (18.2 mg, 91%). Mp 106–107 °C. 87% ee, determined by HPLC [Daicel Chiralcel AD-H column (250 $\times$ 4.6 mm)], n -hexane/ i -PrOH = 60/40, 1.0 mL min $^{-1}$, 254 nm, $t_{\text{major}} = 11.698$ min, $t_{\text{minor}} = 7.095$ min. $[\alpha]_{\text{D}}^{20} = -440.4^\circ$ (c 0.27, CH_2Cl_2). ^1H NMR (400 MHz, DMSO- d_6) δ 9.08 (s, 1H), 8.30–8.24 (m, 1H), 8.11 (s, 1H), 7.77 (d, $J = 7.1$ Hz, 1H), 7.62–7.50 (m, 3H), 7.29 (m, 2H), 7.09 (t, $J = 8.8$ Hz, 2H), 6.60–6.52 (m, 2H), 6.20 (d, $J = 8.6$ Hz, 1H), 3.33 (s, 3H) ppm. ^{13}C NMR (101 MHz, DMSO) δ 168.8, 165.2, 162.8, 160.4, 151.0, 142.0, 132.5, 131.5, 130.3, 129.1, 127.7 (d, $J = 8.0$ Hz), 127.2, 125.5, 123.8, 119.6, 119.3, 119.2, 118.4, 115.4, 115.2, 112.3, 112.0, 66.7, 51.6 ppm. ^{19}F NMR (376 MHz, CDCl_3) δ -116.43 (s) ppm. IR (film): $\gamma = 3403, 2950, 1694, 1601, 1504, 1466, 1437, 1365, 1314, 1267, 1222, 1158, 1093, 1051, 832, 744, 692\text{ cm}^{-1}$. HRMS m/z (ESI^+): calcd for $\text{C}_{24}\text{H}_{17}\text{FN}_2\text{NaO}_3$ ($[\text{M} + \text{Na}]^+$) 423.1115, found 423.1116.

(R)-Methyl 1-(3-oxo-1-(4-(trifluoromethyl)phenyl)isindolin-1-yl)indolizine-2-carboxylate (3h). White solid (18.0 mg, 80%). Mp 110–111 °C. 94% ee, determined by HPLC [Daicel Chiralcel AD-H column (250 $\times$ 4.6 mm)], n -hexane/ i -PrOH = 60/40, 1.0 mL min $^{-1}$, 254 nm, $t_{\text{major}} = 9.671$ min, $t_{\text{minor}} = 5.479$ min. $[\alpha]_{\text{D}}^{20} = -459.6^\circ$ (c 0.30, CH_2Cl_2). ^1H NMR (400 MHz, DMSO- d_6) δ 9.12 (s, 1H), 8.29 (d, $J = 6.6$ Hz, 1H), 8.16 (s, 1H), 7.79 (d, $J = 6.9$ Hz, 1H), 7.64 (d, $J = 8.2$ Hz, 2H), 7.57 (m, 3H), 7.48 (d, $J = 8.1$ Hz, 2H), 6.64–6.52 (m, 2H), 6.14 (d, $J = 9.0$ Hz, 1H), 3.28 (s, 3H) ppm. ^{13}C NMR (101 MHz, CDCl_3) δ 169.3, 165.0, 149.4, 149.2, 131.7, 131.2, 130.4, 129.4, 129.2, 129.1, 125.9, 125.5, 125.3 (d, $J = 3.6$ Hz), 124.7, 122.8, 119.5, 119.4, 119.2, 117.8, 112.3, 112.0, 66.7, 60.9, 29.7, 14.0 ppm. ^{19}F NMR (376 MHz, CDCl_3) δ -62.92 (d, $J = 11.9$ Hz) ppm. IR (film): $\gamma = 3404, 2961, 1694, 1616, 1541, 1503, 1467, 1438, 1411, 1364, 1326, 1263, 1223, 1194, 1163, 1098, 1068, 1017, 802, 757, 744, 695\text{ cm}^{-1}$. HRMS m/z (ESI^+): calcd for $\text{C}_{25}\text{H}_{17}\text{F}_3\text{N}_2\text{NaO}_3$ ($[\text{M} + \text{Na}]^+$) 473.1083, found 473.1086.

Methyl 1-(1-(4-chlorophenyl)-3-oxoisindolin-1-yl)indolizine-2-carboxylate (3i). White solid (16.8 mg, 81%). Mp 126–127 °C. 92% ee, determined by HPLC [Daicel Chiralcel AD-H column (250 $\times$ 4.6 mm)], n -hexane/ i -PrOH = 60/40, 1.0 mL min $^{-1}$, 254 nm, $t_{\text{major}} = 13.252$ min, $t_{\text{minor}} = 7.570$ min. $[\alpha]_{\text{D}}^{20} = -530.6^\circ$ (c 0.39, CH_2Cl_2). ^1H NMR (400 MHz, DMSO- d_6) δ 9.06 (s, 1H), 8.27 (d, $J = 6.2$ Hz, 1H), 8.12 (s, 1H), 7.77 (d, $J = 6.8$ Hz, 1H), 7.60–7.51 (m, 3H), 7.29 (m, 4H), 6.61–6.51 (m, 2H), 6.18 (d, $J = 9.0$ Hz, 1H), 3.33 (s, 3H) ppm. ^{13}C NMR (101 MHz, DMSO) δ 168.8, 165.1, 150.7, 144.9, 132.5, 132.0, 131.4, 130.4, 129.2, 128.7, 128.6, 128.0, 127.5, 127.3, 125.5, 123.9, 119.7, 119.2, 119.1, 118.5, 112.1, 111.9, 66.8, 51.6 ppm. IR (film): $\gamma = 3403, 2950, 1697, 1637, 1488, 1467, 1437, 1364, 1314, 1268, 1221,$

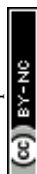
1159, 1093, 1055, 1012, 824, 745, 690 cm^{-1} . HRMS m/z (ESI^+): calcd for $\text{C}_{24}\text{H}_{17}\text{ClN}_2\text{NaO}_3$ ($[\text{M} + \text{Na}]^+$) 439.0820, found 439.0821.

(R)-Methyl 1-(1-(3-chlorophenyl)-3-oxoisindolin-1-yl)indolizine-2-carboxylate (3j). White solid (18.3 mg, 88%). Mp 104–105 °C. 83% ee, determined by HPLC [Daicel Chiralcel AD-H column (250 $\times$ 4.6 mm)], n -hexane/ i -PrOH = 60/40, 1.0 mL min $^{-1}$, 254 nm, $t_{\text{major}} = 9.524$ min, $t_{\text{minor}} = 14.566$ min. $[\alpha]_{\text{D}}^{20} = -521.8^\circ$ (c 0.43, CH_2Cl_2). ^1H NMR (400 MHz, DMSO- d_6) δ 9.11 (s, 1H), 8.28 (d, $J = 6.1$ Hz, 1H), 8.13 (s, 1H), 7.78 (d, $J = 6.8$ Hz, 1H), 7.64–7.51 (m, 3H), 7.28 (t, $J = 4.1$ Hz, 3H), 7.25–7.18 (m, 1H), 6.61–6.52 (m, 2H), 6.18 (d, $J = 8.7$ Hz, 1H), 3.32 (s, 3H) ppm. ^{13}C NMR (101 MHz, DMSO) δ 168.8, 165.1, 150.5, 148.3, 133.4, 132.6, 131.4, 130.6, 130.4, 129.3, 127.4, 127.3, 125.5, 125.4, 124.5, 123.9, 119.7, 119.2, 119.1, 118.5, 112.1, 111.6, 66.9, 51.6 ppm. IR (film): $\gamma = 3398, 2950, 1698, 1611, 1591, 1541, 1503, 1467, 1437, 1370, 1314, 1266, 1221, 1191, 1079, 746, 729, 630\text{ cm}^{-1}$. HRMS m/z (ESI^+): calcd for $\text{C}_{24}\text{H}_{17}\text{ClN}_2\text{NaO}_3$ ($[\text{M} + \text{Na}]^+$) 439.0820, found 439.0823.

(R)-Ethyl 1-(3-oxo-1-phenylisindolin-1-yl)indolizine-2-carboxylate (3k). White solid (16.6 mg, 84%). Mp 115–116 °C. 86% ee, determined by HPLC [Daicel Chiralcel AD-H column (250 $\times$ 4.6 mm)], n -hexane/ i -PrOH = 60/40, 1.0 mL min $^{-1}$, 254 nm, $t_{\text{major}} = 15.741$ min, $t_{\text{minor}} = 9.017$ min. $[\alpha]_{\text{D}}^{20} = -290.1^\circ$ (c 0.91, CH_2Cl_2). ^1H NMR (400 MHz, CDCl_3) δ 7.93 (d, $J = 6.5$ Hz, 1H), 7.88 (s, 1H), 7.80 (d, $J = 9.1$ Hz, 2H), 7.50 (m, 2H), 7.33 (d, $J = 7.1$ Hz, 3H), 7.24–7.16 (m, 3H), 6.44 (t, $J = 6.3$ Hz, 1H), 6.36 (m, 1H), 5.89 (d, $J = 9.5$ Hz, 1H), 4.00 (m, 1H), 3.85 (m, 1H), 1.06 (t, $J = 7.1$ Hz, 3H) ppm. ^{13}C NMR (101 MHz, CDCl_3) δ 169.4, 165.1, 150.3, 145.0, 131.5, 131.2, 130.3, 128.7, 128.2, 127.0, 125.9, 125.7, 125.1, 124.4, 119.9, 119.6, 118.8, 117.5, 112.9, 111.8, 66.9, 60.7, 31.5, 30.2, 14.0 ppm. IR (film): $\gamma = 3403, 2949, 1693, 1610, 1560, 1466, 1437, 1369, 1314, 1267, 1222, 1159, 745, 702\text{ cm}^{-1}$. HRMS m/z (ESI^+): calcd for $\text{C}_{25}\text{H}_{20}\text{N}_2\text{NaO}_3$ ($[\text{M} + \text{Na}]^+$) 419.1366, found 419.1366.

(R)-Ethyl 1-(3-oxo-1-(4-(trifluoromethyl)phenyl)isindolin-1-yl)indolizine-2-carboxylate (3l). White solid (19.3 mg, 83%). Mp 97–99 °C. 94% ee, determined by HPLC [Daicel Chiralcel AD-H column (250 $\times$ 4.6 mm)], n -hexane/ i -PrOH = 60/40, 1.0 mL min $^{-1}$, 254 nm, $t_{\text{major}} = 8.130$ min, $t_{\text{minor}} = 4.912$ min. $[\alpha]_{\text{D}}^{20} = -343.0^\circ$ (c 0.37, CH_2Cl_2). ^1H NMR (400 MHz, CDCl_3) δ 7.96 (d, $J = 7.8$ Hz, 2H), 7.82 (d, $J = 7.7$ Hz, 2H), 7.58–7.45 (m, 6H), 7.31 (d, $J = 7.4$ Hz, 1H), 6.47 (t, $J = 6.7$ Hz, 1H), 6.39 (m, 1H), 5.90 (d, $J = 9.5$ Hz, 1H), 4.02 (m, 1H), 3.96–3.87 (m, 1H), 1.06 (t, $J = 7.1$ Hz, 3H) ppm. ^{13}C NMR (101 MHz, CDCl_3) δ 169.3, 165.0, 149.4, 149.2, 131.7, 131.2, 130.4, 129.4, 129.2, 129.1, 126.8, 125.9, 125.5, 125.3 (d, $J = 3.6$ Hz), 124.7, 122.8, 119.5, 119.4, 119.2, 117.8, 112.3, 112.0, 66.7, 60.9, 29.7, 14.0 ppm. ^{19}F NMR (376 MHz, CDCl_3) δ -62.99 (s) ppm. IR (film): $\gamma = 3403, 2959, 1694, 1615, 1503, 1467, 1326, 1268, 1220, 1163, 1122, 1081, 1017, 837, 744, 695\text{ cm}^{-1}$. HRMS m/z (ESI^+): calcd for $\text{C}_{26}\text{H}_{19}\text{F}_3\text{N}_2\text{NaO}_3$ ($[\text{M} + \text{Na}]^+$) 487.1240, found 487.1241.

(R)-Ethyl 1-(1-(4-chlorophenyl)-3-oxoisindolin-1-yl)indolizine-2-carboxylate (3m). White solid (16.6 mg, 77%). Mp 93–94 °C. 92% ee, determined by HPLC [Daicel Chiralcel AD-H column (250 $\times$ 4.6 mm)], n -hexane/ i -PrOH = 60/40, 1.0 mL min $^{-1}$, 254 nm, $t_{\text{major}} = 13.446$ min, $t_{\text{minor}} = 6.167$ min. $[\alpha]_{\text{D}}^{20} = -507.3^\circ$ (c 0.25, CH_2Cl_2). ^1H NMR (400 MHz, CDCl_3)



δ 7.94 (d, J = 6.9 Hz, 1H), 7.88 (s, 1H), 7.81 (d, J = 4.8 Hz, 2H), 7.52 (m, 2H), 7.32 (d, J = 6.0 Hz, 2H), 7.22 (d, J = 3.7 Hz, 1H), 7.15 (d, J = 4.8 Hz, 2H), 6.45 (t, J = 6.7 Hz, 1H), 6.40–6.34 (m, 1H), 5.87 (d, J = 9.5 Hz, 1H), 4.07 (m, 1H), 3.92 (m, 1H), 1.10 (t, J = 7.1 Hz, 3H) ppm. ^{13}C NMR (101 MHz, CDCl_3) δ 169.3, 165.0, 149.6, 147.3, 134.3, 131.7, 131.2, 130.3, 129.6, 129.1, 127.3, 125.9, 125.8, 125.3, 124.6, 123.4, 119.6, 119.4, 119.1, 117.8, 112.1, 111.9, 66.6, 60.9, 14.1 ppm. IR (film): γ = 3398, 2963, 1694, 1610, 1504, 1466, 1370, 1313, 1267, 1219, 1158, 1096, 830, 745, 692 cm^{-1} . HRMS m/z (ESI^+): calcd for $\text{C}_{25}\text{H}_{19}\text{ClN}_2\text{NaO}_3$ ($[\text{M} + \text{Na}]^+$) 453.0976, found 453.0977.

Methyl 3-(3-oxo-1-phenylisoindolin-1-yl)pyrrolo[1,2-*a*]quinoline-2-carboxylate (3n). White solid (16.8 mg, 78%). Mp 156–157 °C. 91% ee, determined by HPLC [Daicel Chiralcel OD-H column (250 $\times$ 4.6 mm)], *n*-hexane/*i*-PrOH = 85/15, 0.8 mL min^{-1} , 254 nm, t_{major} = 48.470 min, t_{minor} = 34.766 min. $[\alpha]_{\text{D}}^{20}$ = -525.4° (c 0.20, CH_2Cl_2). ^1H NMR (400 MHz, CDCl_3) δ 8.43 (s, 1H), 7.99–7.94 (m, 1H), 7.91–7.85 (m, 2H), 7.58–7.49 (m, 5H), 7.38–7.33 (m, 4H), 7.25–7.19 (m, 2H), 6.72 (d, J = 9.9 Hz, 1H), 5.86 (d, J = 10.0 Hz, 1H), 3.49 (s, 3H) ppm. ^{13}C NMR (101 MHz, CDCl_3) δ 150.1, 144.9, 132.7, 131.7, 131.1, 128.9, 128.9, 128.7, 128.5, 128.4, 127.9, 127.1, 125.9, 125.2, 124.9, 124.5, 123.5, 120.9, 118.8, 118.1, 117.5, 116.4, 114.3, 66.9, 51.6, 31.5, 29.7, 14.2 ppm. IR (film): γ = 3403, 2962, 1694, 1609, 1560, 1513, 1466, 1291, 1261, 1213, 1091, 1030, 799, 752, 701 cm^{-1} . HRMS m/z (ESI^+): calcd for $\text{C}_{28}\text{H}_{20}\text{N}_2\text{NaO}_3$ ($[\text{M} + \text{Na}]^+$) 455.1366, found 455.1367.

(*R*)-Methyl 3-(1-(3-chlorophenyl)-3-oxoisoindolin-1-yl)pyrrolo[1,2-*a*]quinoline-2-carboxylate (3o). White solid (21.2 mg, 91%). Mp 144–145 °C. 87% ee, determined by HPLC [Daicel Chiralcel OD-H column (250 $\times$ 4.6 mm)], *n*-hexane/*i*-PrOH = 85/15, 0.8 mL min^{-1} , 254 nm, t_{major} = 27.451 min, t_{minor} = 23.576 min. $[\alpha]_{\text{D}}^{20}$ = -550.8° (c 0.19, CH_2Cl_2). ^1H NMR (400 MHz, CDCl_3) δ 8.45 (d, J = 5.0 Hz, 1H), 7.96 (t, J = 6.4 Hz, 2H), 7.92–7.88 (m, 1H), 7.59 (d, J = 5.1 Hz, 1H), 7.55 (m, 3H), 7.41–7.38 (m, 1H), 7.34 (d, J = 6.2 Hz, 2H), 7.27 (d, J = 1.9 Hz, 1H), 7.18 (t, J = 4.5 Hz, 2H), 6.73 (m, 1H), 5.84 (m, 1H), 3.57 (m, 3H) ppm. ^{13}C NMR (101 MHz, CDCl_3) δ 169.7, 169.4, 165.2, 149.5, 149.4, 147.1, 142.2, 131.9, 129.9, 129.7, 129.2, 128.8, 128.7, 127.4, 126.0, 125.9, 125.3, 125.1, 124.7, 123.9, 123.5, 122.9, 121.2, 117.9, 114.4, 87.7, 66.6, 51.8 ppm. IR (film): γ = 3403, 2960, 1694, 1610, 1591, 1513, 1438, 1377, 1291, 1261, 1214, 1128, 1088, 794, 750, 699 cm^{-1} . HRMS m/z (ESI^+): calcd for $\text{C}_{28}\text{H}_{19}\text{ClN}_2\text{NaO}_3$ ($[\text{M} + \text{Na}]^+$) 489.0976, found 489.0981.

(*R*)-Methyl 3-(1-(4-chlorophenyl)-3-oxoisoindolin-1-yl)pyrrolo[1,2-*a*]quinoline-2-carboxylate (3p). White solid (15.8 mg, 68%). Mp 156–157 °C. 94% ee, determined by HPLC [Daicel Chiralcel OD-H column (250 $\times$ 4.6 mm)], *n*-hexane/*i*-PrOH = 85/15, 0.8 mL min^{-1} , 254 nm, t_{major} = 31.845 min, t_{minor} = 21.607 min. $[\alpha]_{\text{D}}^{20}$ = -550.8° (c 0.12, CH_2Cl_2). ^1H NMR (400 MHz, CDCl_3) δ 8.44 (s, 1H), 7.98–7.87 (m, 3H), 7.55 (d, J = 7.0 Hz, 4H), 7.37 (t, J = 7.5 Hz, 1H), 7.31 (d, J = 7.5 Hz, 3H), 7.21 (d, J = 8.5 Hz, 2H), 6.73 (d, J = 9.9 Hz, 1H), 5.86 (d, J = 9.9 Hz, 1H), 3.58 (s, 3H) ppm. ^{13}C NMR (151 MHz, CDCl_3) δ 171.9, 167.8, 152.1, 149.7, 136.9, 135.9, 135.2, 134.4, 133.7, 132.2, 131.8, 131.6, 131.4, 131.3, 129.9, 128.5, 127.9, 127.8, 127.3, 126.1, 125.8, 123.7, 120.9, 120.5, 118.3, 116.9, 69.2, 54.3 ppm. IR (film): γ =

3398, 2951, 1694, 1610, 1560, 1514, 1486, 1438, 1359, 1291, 1246, 1213, 1150, 1091, 1014, 793, 752, 692 cm^{-1} . HRMS m/z (ESI^+): calcd for $\text{C}_{28}\text{H}_{19}\text{ClN}_2\text{NaO}_3$ ($[\text{M} + \text{Na}]^+$) 489.0976, found 489.0980.

(*R*)-Methyl 3-(3-oxo-1-(thiophen-2-yl)isoindolin-1-yl)pyrrolo[1,2-*a*]quinoline-2-carboxylate (3q). Yellow solid (13.4 mg, 61%). Mp 144–145 °C. 92% ee, determined by HPLC [Daicel Chiralcel OD-H column (250 $\times$ 4.6 mm)], *n*-hexane/*i*-PrOH = 85/15, 0.8 mL min^{-1} , 254 nm, t_{major} = 38.232 min, t_{minor} = 26.609 min. $[\alpha]_{\text{D}}^{20}$ = -550.8° (c 0.19, CH_2Cl_2). ^1H NMR (400 MHz, CDCl_3) δ 8.41 (s, 1H), 8.01–7.95 (m, 2H), 7.89 (d, J = 8.7 Hz, 1H), 7.58–7.51 (m, 4H), 7.44 (m, 1H), 7.36 (t, J = 7.3 Hz, 1H), 7.11–7.08 (m, 1H), 6.85 (d, J = 3.2 Hz, 2H), 6.72 (d, J = 9.9 Hz, 1H), 5.80 (d, J = 9.9 Hz, 1H), 3.64 (s, 3H) ppm. ^{13}C NMR (101 MHz, CDCl_3) δ 168.9, 165.6, 150.5, 149.8, 132.6, 131.9, 130.6, 129.2, 128.8, 128.7, 128.5, 126.9, 125.7, 125.2, 124.5, 124.2, 123.9, 123.4, 121.1, 119.3, 117.9, 117.3, 115.6, 114.4, 64.6, 51.9 ppm. IR (film): γ = 3403, 2950, 1694, 1601, 1504, 1466, 1437, 1365, 1314, 1267, 1222, 1158, 1093, 1051, 832, 744, 692 cm^{-1} . HRMS m/z (ESI^+): calcd for $\text{C}_{26}\text{H}_{18}\text{N}_2\text{NaO}_3\text{S}$ ($[\text{M} + \text{Na}]^+$) 461.0930, found 461.0931.

Procedure for the 1.0 mmol scale reaction

To a mixture of indolizine-2-carboxylate **1c** (225 mg, 1 mmol), 3-hydroxyisoindolinones **2i** (259 mg, 1 mmol, 1 equiv.) and catalyst (*S*)-**4a** (0.1 mmol, 10 mol%) in DCE (20 mL) was added 4 Å MS (2 g). After stirring at room temperature for 24 hours, the residue was purified by flash column chromatography with acetate/petroleum ether 1 : 2 (v/v) on silica gel to give the desired product **3p** (284 mg) in 61% yield with 94% ee.

Procedure for the derivatization experiment

To a 100 mL round-bottomed flask, **3p** (292 mg, 0.63 mmol), NaOH (125 mg, 3.2 mmol) and 30 mL mixed solvent [THF/MeOH/ H_2O 2 : 2 : 0.5 (v/v/v)] were added. After refluxing for 5 hours, adjusted the pH to 2 with 1 M HCl and extracted three times with ethyl acetate. The residue was purified by flash column chromatography with acetate/petroleum ether/formic acid 1 : 1 : 1.5% (v/v/v) on silica gel to give the desired product **4** (230 mg) in 81% yield with 98% ee.

(*R*)-3-(1-(4-Chlorophenyl)-3-oxoisoindolin-1-yl) pyrrolo[1,2-*a*]quinoline-2-carboxylic acid. Yellow solid (230 mg, 81%). Mp 212–213 °C. 98% ee, determined by HPLC [Daicel AD-H column (250 $\times$ 4.6 mm)], *n*-hexane/*i*-PrOH/ HCOOH = 80/20/0.1, 1.0 mL min^{-1} , 254 nm, t_{major} = 21.848 min, t_{minor} = 12.595 min. $[\alpha]_{\text{D}}^{20}$ = -568.889° (c 0.13, CH_2Cl_2). ^1H NMR (400 MHz, CD_2Cl_2) δ 13.31 (s, 1H), 10.37 (s, 1H), 8.56 (s, 1H), 7.92 (m, 2H), 7.59 (m, 2H), 7.53 (m, 2H), 7.47–7.43 (m, 1H), 7.38–7.24 (m, 5H), 6.74 (d, J = 10.0 Hz, 1H), 6.10 (d, J = 10.0 Hz, 1H) ppm. ^{13}C NMR (101 MHz, CD_2Cl_2) δ 171.5, 167.9, 150.1, 144.1, 133.13, 133.09, 132.2, 131.6, 129.7, 129.5, 129.1, 128.9, 128.5, 127.0, 126.8, 125.5, 124.7, 123.8, 121.2, 119.6, 118.4, 116.6, 114.8, 68.4, 54.4, 54.1, 53.8, 53.5, 53.3 ppm. IR (film): γ = 3290, 2924, 1689, 1661, 1610, 1540, 1488, 1441, 1379, 1329, 1290, 1213, 1154, 1014, 918, 828, 791, 750, 704 cm^{-1} . HRMS m/z (ESI^+): calcd for $\text{C}_{27}\text{H}_{17}\text{ClN}_2\text{NaO}_3$ ($[\text{M} + \text{Na}]^+$) 475.0820, found 475.0820.



Conflicts of interest

There are no conflicts to declare.

Acknowledgements

This work was supported by the National Natural Science Foundation of China (22071213), Leading Talents of Special Support Program of Zhejiang Province High-level Talents (2020R52008), the Fundamental Research Funds for the Central Universities (226-2022-00224) and Center of Chemistry for Frontier Technologies of Zhejiang University.

Notes and references

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