

Synthesis of Dihydropyrrole Derivatives by Oxidative Functionalization of 2-Amino-4*H*-Chromenes Using Hypervalent Iodine Reagents

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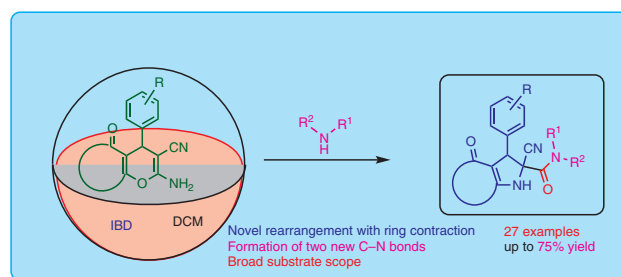
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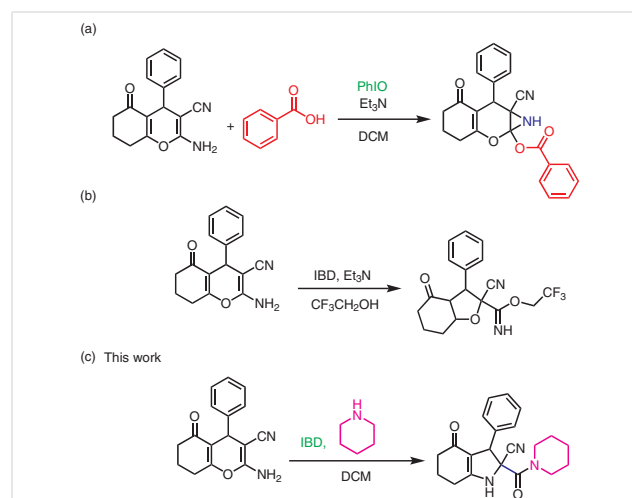
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Abstract An efficient simple, metal-free, one-pot protocol for the synthesis of dihydropyrrole derivatives has been achieved via sequential addition of iodobenzenediacetate and secondary amine to 2-amino-4*H*-pyran derivatives. The one-pot protocol proceeds through tandem oxidative functionalization, rearrangement, and ring contraction to provide an entirely new strategy for the construction of the dihydropyrrole skeleton.

Key words one pot, metal free, oxidative functionalization, rearrangement, ring contraction

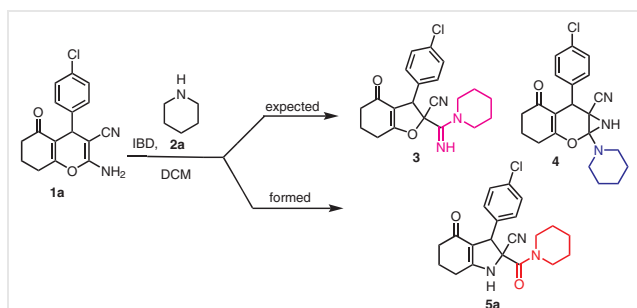
Hypervalent iodine (HVI) reagents have fascinated chemists over many years as versatile synthetic agents. HVIs are mildly oxidizing reagents, possessing electrophilic character, environmentally friendly behavior, and are commercially accessible.^{1,2} Occasionally, they are found to be superior agents than metal catalysts.³ HVI reagents mediate several group transformations on various scaffolds involving oxidative functionalization,⁴ intermolecular rearrangements,⁵ and cyclization/coupling reactions.^{6,7} They have been known to be helpful in aminofluorination,⁸ diamination,⁹ dioxygenation,¹⁰ halogenations,¹¹ 1,5-electrocyclization,¹² and acetoxylation.¹³ The electrophilic and excellent ligand-exchange¹⁴ nature of the iodine in HVI reagents makes them suitable for the generation of the cationic intermediates which can react with nucleophiles or form rearranged products with ring expansion, ring contraction, or migration of functional groups.¹⁵ The geminal dialkoxylation and 1,2-migration of $-NH_2$ in 2-amino-4*H*-pyrans was reported by this group in the presence of iodobenzene diacetate (IBD) via an apparent intramolecular aziridination.^{16a} The same study with *N*-chlorosuccinimide (NCS) shows

both chlorination and alkoxylation at the double bond in the 2-amino-4*H*-pyrans.^{16a} Yet another oxidative difunctionalization of amino pyrans leading to dihydrofurans through sequential addition of NCS in the presence of base has also been accomplished by this group.^{16b} Zhao et al., demonstrated the conventional method for the conversion of enamines to 2-*H*-aziridines and their subsequent rearrangements by using IBD.¹⁷ Recently, Das et al. have reported a synthetic route to construct the aziridine ring by exploiting the enamine fragment of 2-amino-4*H*-pyrans and their skeletal transformation in the presence of IBD (Scheme 1a).¹⁸ Synthesis of 2,2,2-trifluoroethyl 2-cyano-4-oxo-3-phenyloctahydrobenzofuran-2-carbimide from 2-amino-4*H*-pyrans has also been accomplished (Scheme 1b).^{19a} The syntheses of dihydrofuran,^{16b,19} pyridone,²⁰ aziridine,²¹ and oxazine²² derivatives through the rearrangement of 2-amino-4*H*-pyrans have also been accomplished.



Scheme 1 Previous and present approaches towards the oxidative functionalization of amino chromenes

Fused heterocycles are formed by the union of two or more heterocyclic frameworks into a single molecular identity. They are found in many natural products which exhibit a broad variety of biological activities primarily owing to the presence of multiple pharmacophores in a single molecular entity.²³ Among these, fused pyrrole heterocycles are found in large variety of naturally occurring compounds.²⁴ Inherent diversity of dihydropyrrole derivatives and their distinct therapeutic response led to many researchers choosing them for exploring their maximum potential as medicinal and pharmaceutical agents,²⁵ as well as in materials science.²⁶ Owing to the importance of dihydropyrrole scaffolds, synthesis and/or derivatization of these compounds is of contemporary interest. Keeping this in consideration and as a part of our ongoing research program on the development of innovative and efficient synthetic protocols for the construction of biologically significant heterocycles, herein a report on the synthesis of functionalized dihydropyrroles through skeletal transformation of 2-amino chromenes is described for the first time under one-pot conditions mediated by HVI (Scheme 2).

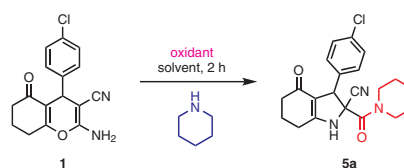


Scheme 2 Synthesis of compound 5

2-Amino-4-(4-chlorophenyl)-5-oxo-5,6,7,8-tetrahydro-4H-chromene-3-carbonitrile was synthesized from a method previously developed in our laboratory.¹⁶ In an effort towards oxidative functionalization of 2-amino chromenes employing 2-amino-4-(4-chlorophenyl)-5-oxo-5,6,7,8-tetrahydro-4H-chromene-3-carbonitrile as model substrate the reaction with IBD (1 mmol) has been investigated in 4 mL of dichloromethane (DCM) at room temperature (rt) for 30 min, followed by addition of piperidine (1 mmol) to the reaction mixture. The complete consumption of the starting materials was monitored by TLC, and column chromatography of the reaction mixture gave white-colored solid in 65% yield (Table 1, entry 1). An anticipated structure of the product can be either compound **3** or **4**. Normally, imines on hydrolysis under acidic conditions result in the formation of the corresponding carbonyl compound.^{16b} To confirm the structure of the compound, acidic hydrolysis of the product was performed. No change in the TLC and mass spectrum of the product was observed. Therefore structure **3** was ruled out. The ¹H NMR spectrum of the compound shows a singlet signal at $\delta = 5.49$ ppm accounting for $-\text{NH}$,

and the ¹³C NMR spectra of compound a signal at $\delta = 164$ ppm, likely due to the presence of C=O in the formed compound, firmly ruling out compound **4** as the product. Ultimately, the structure of the compound was confirmed by X-ray crystallographic studies and assigned as 3-(4-chlorophenyl)-4-oxo-2-(piperidine-1-carbonyl)-2,3,4,5,6,7-hexahydro-1H-indole-2-carbonitrile (**5a**).

Table 1 Optimization of Reaction Conditions for the Synthesis of 3-(4-Chlorophenyl)-4-oxo-2-(piperidine-1-carbonyl)-2,3,4,5,6,7-hexahydro-1H-indole-2-carbonitrile (**5a**)^a



Entry	Solvent	Oxidant	Temp	Time (h)	Yield (%) ^b
1	DCM	IBD	rt	6	65
2	MeCN	IBD	rt	6	50
3	toluene	IBD	rt	6	45
4	DCE	IBD	rt	6	52
5	THF	IBD	rt	6	traces
6	dioxane	IBD	rt	6	30
7	DMF	IBD	rt	6	NR ^c
8	H ₂ O	IBD	rt	6	NR ^c
9	MeOH	IBD	rt	1	NR ^c
10	DCM	IBD (1.1) ^d	rt	6	70
11	DCM	IBD (1.2) ^d	rt	6	59
12	DCM	IBD (1.5) ^d	rt	6	50
13	DCM	IBD (1.1) ^d	0 °C	8	30
13	DCM	PIFA	rt	6	traces
14	DCM	HTIB	rt	6	NR ^c
15	DCM	PhIO	rt	6	NR ^c

^a Reaction conditions: 2-amino-4-(4-chlorophenyl)-5-oxo-5,6,7,8-tetrahydro-4H-chromene-3-carbonitrile (1 mmol), IBD (1 mmol), piperidine (1 mmol).

^b Isolated yields.

^c NR: no reaction.

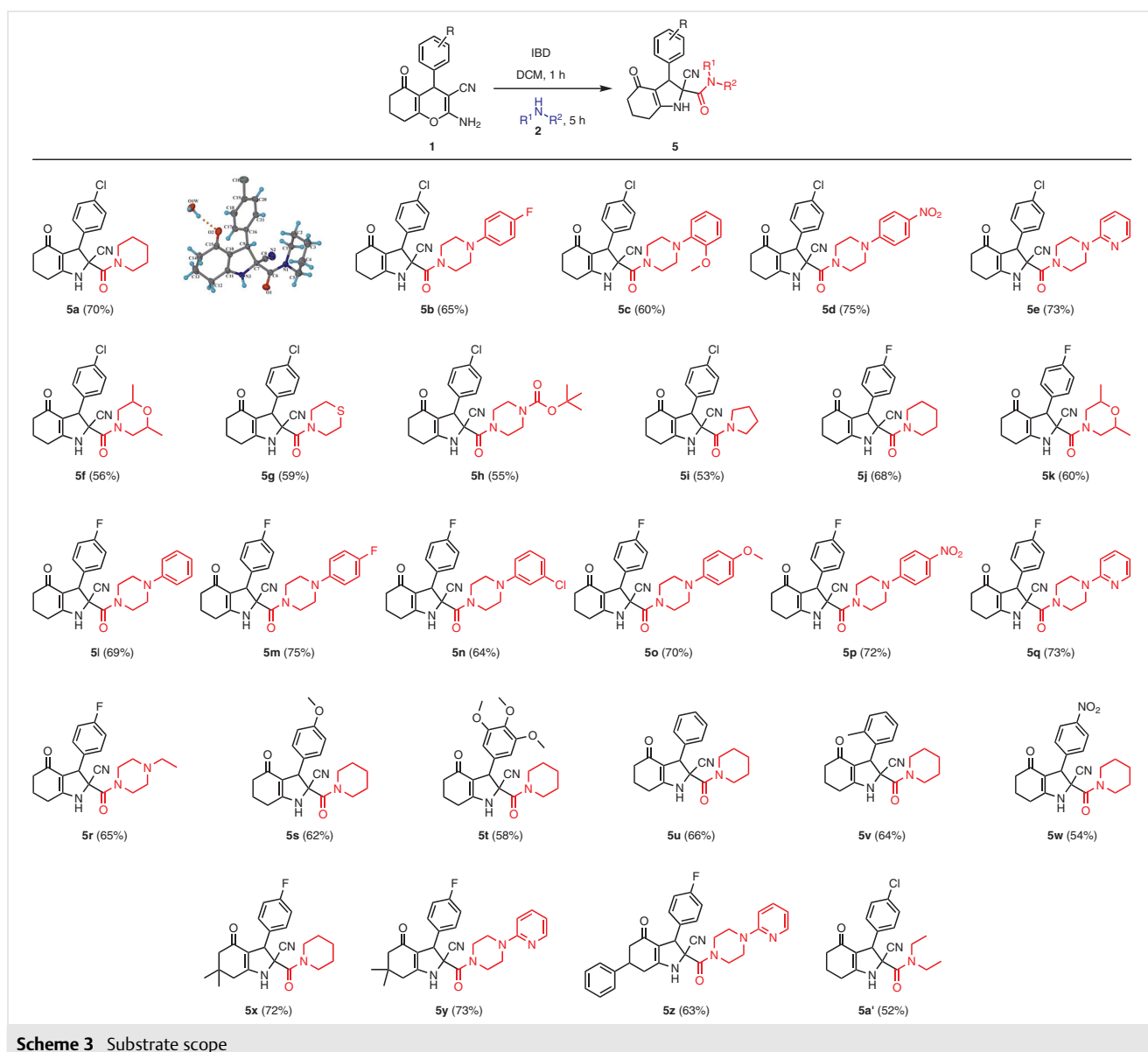
^d Equivalents of oxidant used in parentheses.

After the conformation of the product structure, work on identification of suitable reaction conditions for the synthesis of 3-(4-chlorophenyl)-4-oxo-2-(piperidine-1-carbonyl)-2,3,4,5,6,7-hexahydro-1H-indole-2-carbonitrile was initiated by altering various reaction parameters (Table 1). Firstly, the solvent effect was studied by changing solvents such as acetonitrile (MeCN), toluene, dichloroethane (DCE), tetrahydrofuran (THF), and dioxane. The reaction time as well as yield of the reaction and product distribution was greatly affected in all the cases and a decrease in the product yield was observed (Table 1, entries 2–6). In di-

methylformamide (DMF) and water (H₂O), formation of the product **5** was not observed (Table 1, entries 7 and 8). Whereas in methanol the formation of 3-amino-4-(4-chlorophenyl)-2,2-dimethoxy-5-oxo-3,4,5,6,7,8-hexahydro-2H-chromene-3-carbonitrile was observed, as mentioned in an earlier report^{16a} (Table 1, entry 9). Alternative HVI reagents were studied. With iodosobenzene bis(trifluoroacetate) (PIFA, Table 1, entry 13) only traces of compound were formed, whereas with other reagents, HTIB and PhIO (hydroxyl (tosyloxy)iodobenzene and iodosobenzene, Table 1, entries 14 and 15), no product formation was observed (starting material recovered). Later ideal HVI load was determined and by changing the quantity of IBD. Raising the molar ratio to 1.1 equivalents in solvent DCM increased the

product yield to 70% (Table 1 entry 10). Further increase in the mol equivalents of IBD (Table 1, entries 11 and 12) did not significantly increase the product yield.

After establishing the feasible reaction conditions for the preparation of compound **5**, an exploration of the substrate scope was taken up, using a variety of secondary amines as exemplified in Scheme 3. Secondary amines such as 1-phenylpiperazine derivatives with neutral, electron-donating (-OMe) and electron-withdrawing group (-NO₂) on the phenyl ring gave moderate to good yields (**5b–e,h**). Morpholine and thiomorpholine derivatives afforded moderate yields (**5f,g**). Similarly, the reaction with pyrrole afforded the product **5i** in moderate yield. Subsequently the substrate scope was extended to various neutral, electron-



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- (27) **General Procedure for the Preparation of Dihydropyrrole Derivatives**
A mixture of 2-amino-5-oxo-4-phenyl-5,6,7,8-tetrahydro-4H-chromene-3-carbonitrile (1 mmol, 1 equiv) and IBD (1.1 mmol, 1.1 equiv) in anhydrous DCM (3 mL) was stirred for 60 min, then secondary amine derivatives were added. Stirring was continued at room temperature until the starting material was completely consumed (TLC monitoring). After completion, the reaction mixture was extracted with DCM (20 mL) and washed with water (10 mL). The combined organic layers were dried (anhydrous Na₂SO₄) and evaporated under reduced pressure to dryness. The crude product thus obtained was purified by column chromatography (60–120 mesh, and EtOAc-hexane, 30:70) to afford the pure product.
- (28) **Analytical Data**
3-(4-Chlorophenyl)-4-oxo-2-(piperidine-1-carbonyl)-2,3,4,5,6,7-hexahydro-1H-indole-2-carbonitrile (5a)
Yield: 134 mg (70%); white solid; mp 156–158 °C. ¹H NMR (500 MHz, CDCl₃): δ = 7.35 (d, J = 8.4 Hz, 2 H), 7.25 (dd, J = 7.8, 5.4 Hz, 2 H), 5.53 (s, 1 H), 4.39 (s, 1 H), 3.58 (m, 4 H), 2.59 (q, J = 6.4 Hz, 2 H), 2.29–2.22 (m, 2 H), 2.11–2.03 (m, 2 H), 1.68–1.58 (m, 2 H), 1.30 (dt, J = 13.8, 7.0 Hz, 2 H), 0.88 (t, J = 7.1 Hz, 2 H). ¹³C NMR (101 MHz, CDCl₃): δ = 192.38, 167.41, 163.51, 136.17, 134.48, 129.36, 115.88, 77.06, 67.33, 54.35, 47.59, 45.53, 36.44, 25.36, 25.13, 24.06, 23.70, 22.20. HRMS-ESI: m/z [M + H]⁺ calcd for C₂₁H₂₅O₂N₃Cl: 384.14869; found: 384.14733.
- 3-(4-Chlorophenyl)-2-[4-(2-fluorophenyl)piperazine-1-carbonyl]-4-oxo-2,3,4,5,6,7-hexahydro-1H-indole-2-carbonitrile (5b)**
Yield: 155 mg (65%); white solid; mp 212–214 °C. ¹H NMR (500 MHz, CDCl₃): δ = 7.39–7.36 (m, 2 H), 7.28–7.25 (m, 2 H), 7.02–6.97 (m, 2 H), 6.92–6.88 (m, 2 H), 5.47 (s, 1 H), 4.44 (s, 1 H), 4.07–3.62 (m, 4 H), 3.31–3.01 (m, 4 H), 2.61 (dd, J = 15.8, 6.7 Hz, 2 H), 2.29–2.24 (m, 2 H), 2.10–2.07 (m, 2 H). ¹³C NMR (101 MHz, CDCl₃): δ = 192.40, 167.21, 163.92, 159.26, 156.80, 147.11, 135.86, 134.68, 129.40, 119.02, 115.99, 115.77, 67.28, 54.55, 50.40, 49.76, 46.59, 44.22, 36.43, 23.70, 22.20. HRMS-ESI: m/z [M + H]⁺ calcd for C₂₆H₂₅O₂N₄ClF: 479.16507; found: 479.16446.
- 3-(4-Chlorophenyl)-2-[4-(2-methoxyphenyl)piperazine-1-carbonyl]-4-oxo-2,3,4,5,6,7-hexahydro-1H-indole-2-carbonitrile (5c)**
Yield: 147 mg (60%); off-white solid; mp 204–206 °C. ¹H NMR (500 MHz, CDCl₃): δ = 7.32 (dd, J = 5.4, 3.5 Hz, 2 H), 7.29–7.27 (m, 1 H), 7.21 (dt, J = 6.8, 1.9 Hz, 1 H), 7.06 (m, 1 H), 6.95–6.89 (m, 3 H), 5.51 (s, 1 H), 4.44 (s, 1 H), 3.95 (s, 1 H), 3.90 (s, 3 H),

3.87–3.75 (m, 3 H), 3.22 (dd, $J = 10.3, 5.9$ Hz, 3 H), 3.07–2.99 (m, 1 H), 2.66–2.57 (m, 2 H), 2.31–2.26 (m, 2 H), 2.12–2.08 (m, 2 H). ^{13}C NMR (101 MHz, CDCl_3): $\delta = 192.36, 167.51, 163.83, 152.30, 139.97, 139.42, 135.02, 130.44, 129.02, 128.12, 126.37, 124.04, 121.16, 118.72, 115.76, 115.55, 111.46, 67.26, 55.55, 54.73, 50.24, 49.80, 46.98, 44.54, 36.44, 23.71, 22.18$. HRMS-ESI: m/z [$M + H$] $^+$ calcd for $\text{C}_{27}\text{H}_{28}\text{O}_3\text{N}_4\text{Cl}$: 491.18498; found: 491.18444.

3-(4-Chlorophenyl)-2-[4-(4-nitrophenyl)piperazine-1-carbonyl]-4-oxo-2,3,4,5,6,7-hexahydro-1H-indole-2-carbonitrile (5d)

Yield: 189 mg (75%); yellow solid; mp 246–248 °C. ^1H NMR (500 MHz, CDCl_3): $\delta = 8.18$ – 8.16 (m, 1 H), 7.35 (dd, $J = 4.7, 1.9$ Hz, 1 H), 7.31–7.28 (m, 1 H), 7.24–7.21 (m, 1 H), 6.91–6.87 (m, 2 H), 5.42 (s, 1 H), 4.42 (s, 1 H), 4.20–4.07 (m, 1 H), 3.93 (s, 1 H), 3.70 (dd, $J = 68.9, 29.8$ Hz, 4 H), 3.49–3.31 (m, 2 H), 2.70–2.57 (m, 2 H), 2.31–2.26 (m, 2 H), 2.14–2.07 (m, 2 H). ^{13}C NMR (101 MHz, $\text{CDCl}_3 + \text{DMSO}$): $\delta = 189.86, 167.29, 162.23, 153.13, 139.11, 137.40, 132.97, 129.03, 127.10, 126.91, 125.60, 124.51, 114.81, 112.00, 110.84, 66.83, 52.69, 44.93, 35.20, 22.22, 20.96$. HRMS-ESI: m/z [$M + H$] $^+$ calcd for $\text{C}_{26}\text{H}_{25}\text{O}_4\text{N}_5\text{Cl}$: 506.16003; found: 506.15896

3-(4-Chlorophenyl)-4-oxo-2-[4-(pyridin-2-yl)piperazine-1-carbonyl]-2,3,4,5,6,7-hexahydro-1H-indole-2-carbonitrile (5e)

Yield: 168 mg (73%); white solid; mp 204–206 °C. ^1H NMR (400 MHz, CDCl_3): $\delta = 8.22$ (ddd, $J = 4.9, 1.9, 0.8$ Hz, 1 H), 7.57–7.50 (m, 1 H), 7.42–7.33 (m, 1 H), 7.29–7.24 (m, 2 H), 6.75–6.66 (m, 2 H), 5.52 (s, 1 H), 4.44 (s, 1 H), 4.01–3.41 (m, 8 H), 2.60 (m, 2 H), 2.30–2.24 (m, 2 H), 2.12–2.06 (m, 2 H). ^{13}C NMR (101 MHz, CDCl_3): $\delta = 192.36, 167.27, 164.09, 158.70, 148.09, 137.89, 135.88, 134.66, 129.43, 115.84, 115.62, 114.57, 107.45, 77.06, 67.38, 54.51, 46.22, 44.95, 44.49, 43.98, 36.42, 23.69, 22.20$. HRMS-ESI: m/z [$M + H$] $^+$ calcd for $\text{C}_{25}\text{H}_{25}\text{O}_2\text{N}_5\text{Cl}$: 462.17025; found: 462.16913.

3-(4-Chlorophenyl)-2-(2,6-dimethylmorpholine-4-carbonyl)-4-oxo-2,3,4,5,6,7-hexahydro-1H-indole-2-carbonitrile (5f)

Yield: 116 mg (56%); off-white solid; mp 200–202 °C. ^1H NMR (400 MHz, CDCl_3): $\delta = 7.37$ – 7.27 (m, 2 H), 7.18 (dd, $J = 36.9, 6.9$ Hz, 2 H), 5.74 (s, 1 H), 4.34 (dd, $J = 23.1, 13.8$ Hz, 2 H), 3.81 (d, $J = 12.8$ Hz, 1 H), 3.67 (s, 2 H), 3.09–2.82 (m, 1 H), 2.59 (d, $J = 27.8$ Hz, 3 H), 2.26 (d, $J = 6.3$ Hz, 2 H), 2.13–2.01 (m, 2 H), 1.33–1.13 (m, 6 H). ^{13}C NMR (101 MHz, CDCl_3): $\delta = 192.33, 167.63, 163.92, 139.24, 134.97, 130.41, 129.07, 128.14, 126.26, 115.93, 115.21, 71.76, 71.12, 70.57, 67.35, 55.68, 53.83, 52.14, 51.79, 49.39, 48.95, 36.39, 23.65, 22.16, 18.65, 18.45$. HRMS-ESI: m/z [$M + H$] $^+$ calcd for $\text{C}_{22}\text{H}_{25}\text{O}_3\text{N}_3\text{Cl}$: 414.15821; found: 414.15790.

3-(4-Chlorophenyl)-4-oxo-2-(thiomorpholine-4-carbonyl)-2,3,4,5,6,7-hexahydro-1H-indole-2-carbonitrile (5g)

Yield: 118 mg (59%); white solid; mp 214–216 °C. ^1H NMR (500 MHz, CDCl_3): $\delta = 7.32$ (td, $J = 4.6, 3.0$ Hz, 2 H), 7.28–7.25 (m, 1 H), 7.21–7.17 (m, 1 H), 5.42 (s, 1 H), 4.35 (s, 1 H), 4.31 (d, $J = 11.7$ Hz, 1 H), 4.02 (d, $J = 10.2$ Hz, 1 H), 3.80–3.71 (m, 1 H), 3.64–3.53 (m, 1 H), 2.96–2.85 (m, 1 H), 2.80–2.71 (m, 1 H), 2.61 (m, 4 H), 2.28 (dd, $J = 9.5, 4.0$ Hz, 2 H), 2.12–2.05 (m, 2 H). ^{13}C NMR (101 MHz, CDCl_3): $\delta = 192.37, 167.46, 164.06, 139.27, 135.07, 130.49, 129.12, 128.06, 126.27, 115.71, 67.36, 54.60, 48.96, 47.03, 36.40, 27.22, 26.65, 23.69, 22.17$. HRMS-ESI: m/z [$M + H$] $^+$ calcd for $\text{C}_{20}\text{H}_{21}\text{O}_2\text{N}_3\text{ClS}$: 402.10532; found: 402.10509.

tert-Butyl 4-[3-(4-Chlorophenyl)-2-cyano-4-oxo-2,3,4,5,6,7-hexahydro-1H-indole-2-carbonyl]piperazine-1-carboxylate (5h)

Yield: 134 mg (55%); white solid; mp 118–120 °C. ^1H NMR (500

MHz, CDCl_3): $\delta = 7.37$ (d, $J = 8.4$ Hz, 2 H), 7.24 (d, $J = 8.4$ Hz, 2 H), 5.45 (s, 1 H), 4.38 (s, 1 H), 3.96–3.25 (m, 8 H), 2.65–2.56 (m, 2 H), 2.29–2.24 (m, 2 H), 2.11–2.04 (m, 2 H), 1.48 (s, 9 H). ^{13}C NMR (101 MHz, CDCl_3): $\delta = 192.40, 167.21, 164.17, 154.36, 135.76, 134.70, 129.50, 115.84, 115.65, 80.92, 77.06, 67.29, 54.48, 46.36, 44.13, 36.41, 28.37, 23.68, 22.18$. HRMS-ESI: m/z [$M + H$] $^+$ calcd for $\text{C}_{25}\text{H}_{30}\text{O}_4\text{N}_4\text{Cl}$: 485.19561; found: 485.19501.

3-(4-Chlorophenyl)-4-oxo-2-(pyrrolidine-1-carbonyl)-2,3,4,5,6,7-hexahydro-1H-indole-2-carbonitrile (5i)

Yield: 98 mg (53%); off-white solid; mp 258–260 °C. ^1H NMR (400 MHz, CDCl_3): $\delta = 7.35$ (d, $J = 8.4$ Hz, 2 H), 7.25 (d, $J = 8.6$ Hz, 2 H), 5.47 (s, 1 H), 4.47 (s, 1 H), 3.75–3.40 (m, 4 H), 2.60 (t, $J = 5.6$ Hz, 2 H), 2.37–2.21 (m, 2 H), 2.12–2.02 (m, 4 H), 1.96 (dd, $J = 13.1, 5.7$ Hz, 2 H). ^{13}C NMR (101 MHz, CDCl_3): $\delta = 192.58, 167.69, 162.77, 136.22, 134.39, 129.30, 115.41, 77.06, 67.97, 53.43, 48.51, 47.56, 36.43, 26.70, 23.65, 23.34, 22.22$. HRMS-ESI: m/z [$M + H$] $^+$ calcd for $\text{C}_{20}\text{H}_{21}\text{O}_2\text{N}_3\text{Cl}$: 370.13359; found: 370.13302.

3-(4-Fluorophenyl)-4-oxo-2-(piperidine-1-carbonyl)-2,3,4,5,6,7-hexahydro-1H-indole-2-carbonitrile (5j)

Yield: 98 mg (73%); white solid; mp 210–212 °C. ^1H NMR (400 MHz, CDCl_3): $\delta = 7.36$ – 7.12 (m, 2 H), 7.14–6.90 (m, 2 H), 5.44 (s, 1 H), 4.40 (s, 1 H), 3.84–3.36 (m, 4 H), 2.67–2.46 (m, 2 H), 2.34–2.17 (m, 2 H), 2.09 (dd, $J = 12.3, 6.1$ Hz, 2 H), 1.86–1.58 (m, 6 H). ^{13}C NMR (101 MHz, CDCl_3): $\delta = 192.36, 167.35, 163.89, 163.58, 161.43, 133.46, 129.65, 116.18, 115.96, 115.76, 67.51, 54.19, 47.56, 45.47, 36.43, 25.34, 25.11, 24.04, 23.66, 22.19$. HRMS-ESI: m/z [$M + H$] $^+$ calcd for $\text{C}_{20}\text{H}_{21}\text{O}_2\text{N}_3\text{Cl}$: 354.17359; found: 354.17302

2-(2,6-Dimethylmorpholine-4-carbonyl)-3-(4-fluorophenyl)-4-oxo-2,3,4,5,6,7-hexahydro-1H-indole-2-carbonitrile (5k)

Yield: 119 mg (60%); off-white solid; mp 150–152 °C. ^1H NMR (500 MHz, CDCl_3): $\delta = 7.26$ (t, $J = 10.3$ Hz, 2 H), 7.08 (t, $J = 8.4$ Hz, 2 H), 5.49 (s, 1 H), 4.47–4.24 (m, 2 H), 4.06 (dd, $J = 83.5, 21.9$ Hz, 1 H), 3.82 (d, $J = 12.4$ Hz, 1 H), 3.64 (d, $J = 31.2$ Hz, 2 H), 2.58 (tt, $J = 23.9, 11.8$ Hz, 3 H), 2.29 (t, $J = 19.2$ Hz, 2 H), 2.15–2.00 (m, 2 H), 1.21 (dd, $J = 47.3, 12.0$ Hz, 6 H). ^{13}C NMR (101 MHz, CDCl_3): $\delta = 192.47, 167.16, 164.01, 163.66, 133.14, 129.71, 116.34, 116.12, 71.81, 71.09, 70.64, 67.43, 55.45, 53.72, 52.16, 49.43, 36.45, 23.71, 22.21, 18.67$. HRMS-ESI: m/z [$M + H$] $^+$ calcd for $\text{C}_{22}\text{H}_{25}\text{O}_3\text{N}_3\text{F}$: 398.18865; found: 398.18845.

3-(4-Fluorophenyl)-4-oxo-2-(4-phenylpiperazine-1-carbonyl)-2,3,4,5,6,7-hexahydro-1H-indole-2-carbonitrile (5l)

Yield: 153 mg (69%); white solid; mp 232–234 °C. ^1H NMR (500 MHz, CDCl_3): $\delta = 7.32$ – 7.27 (m, 1 H), 7.12–7.07 (m, 1 H), 6.95 (dd, $J = 7.7, 3.6$ Hz, 1 H), 5.49 (s, 1 H), 4.47 (s, 1 H), 4.06–3.58 (m, 4 H), 3.48–2.89 (m, 4 H), 2.70–2.49 (m, 2 H), 2.42–2.19 (m, 2 H), 2.09 (t, $J = 6.3$ Hz, 2 H). ^{13}C NMR (101 MHz, CDCl_3): $\delta = 192.42, 167.18, 164.01, 161.54, 150.48, 133.24, 129.74, 129.37, 121.23, 117.05, 116.36, 116.14, 115.94, 67.49, 54.43, 49.41, 48.82, 46.54, 44.19, 36.45, 23.69, 22.21$. HRMS-ESI: m/z [$M + H$] $^+$ calcd for $\text{C}_{26}\text{H}_{26}\text{O}_2\text{N}_4\text{F}$: 445.20385; found: 445.20343.

3-(4-Fluorophenyl)-2-[4-(4-fluorophenyl)piperazine-1-carbonyl]-4-oxo-2,3,4,5,6,7-hexahydro-1H-indole-2-carbonitrile (5m)

Yield: 173 mg (75%); white solid; mp 230–232 °C. ^1H NMR (500 MHz, CDCl_3): $\delta = 7.34$ – 7.27 (m, 2 H), 7.12–7.05 (m, 2 H), 7.03–6.96 (m, 2 H), 6.93–6.84 (m, 2 H), 5.46 (s, 1 H), 4.46 (s, 1 H), 4.03 (d, $J = 12.5$ Hz, 1 H), 3.90–3.57 (m, 3 H), 3.35–2.99 (m, 4 H), 2.60 (m, 2 H), 2.31–2.24 (m, 2 H), 2.09 (t, $J = 6.2$ Hz, 2 H). ^{13}C NMR (101 MHz, CDCl_3): $\delta = 192.45, 167.07, 164.03, 161.57, 159.22, 156.82, 147.11, 133.20, 129.81, 129.72, 119.04, 116.39, 116.18,$

116.00, 115.78, 67.45, 54.47, 50.42, 49.79, 46.60, 44.22, 36.47, 23.73, 22.23. HRMS-ESI: m/z [M + H]⁺ calcd for C₂₆H₂₅O₂N₄F₂: 463.19430; found: 463.19401.

2-[4-(3-Chlorophenyl)piperazine-1-carbonyl]-3-(4-fluorophenyl)-4-oxo-2,3,4,5,6,7-hexahydro-1H-indole-2-carbonitrile (5n)

Yield: 134 mg (64%); white solid; mp 248–250 °C. ¹H NMR (500 MHz, CDCl₃): δ = 7.32–7.28 (m, 2 H), 7.22–7.18 (m, 1 H), 7.12–7.07 (m, 2 H), 6.92–6.89 (m, 2 H), 6.82–6.78 (m, 1 H), 5.44 (s, 1 H), 4.45 (s, 1 H), 4.13–3.57 (m, 4 H), 3.24 (dd, J = 71.6, 62.7 Hz, 4 H), 2.65–2.57 (m, 2 H), 2.28 (m, 2 H), 2.12–2.07 (m, 2 H). ¹³C NMR (101 MHz, CDCl₃): δ = 192.44, 167.13, 164.05, 161.56, 151.51, 135.17, 133.15, 130.30, 129.80, 129.72, 120.90, 116.89, 116.40, 116.18, 115.90, 115.84, 114.80, 77.06, 67.47, 54.42, 48.88, 48.30, 46.31, 43.99, 36.44, 23.70, 22.21. HRMS-ESI: m/z [M + H]⁺ calcd for C₂₆H₂₅O₂N₄ClF: 479.16507; found: 479.16446.

3-(4-Fluorophenyl)-2-[4-(4-methoxyphenyl)piperazine-1-carbonyl]-4-oxo-2,3,4,5,6,7-hexahydro-1H-indole-2-carbonitrile (5o)

Yield: 166 mg (70%); off-white solid; mp 158–160 °C. ¹H NMR (500 MHz, CDCl₃): δ = 7.32–7.28 (m, 2 H), 7.11–7.06 (m, 2 H), 6.94–6.89 (m, 2 H), 6.87–6.83 (m, 2 H), 5.47 (s, 1 H), 4.46 (s, 1 H), 4.00 (d, J = 11.1 Hz, 1 H), 3.83 (s, 1 H), 3.78 (s, 3 H), 3.76–3.64 (m, 2 H), 3.27–2.97 (m, 4 H), 2.61 (q, J = 6.5 Hz, 2 H), 2.27 (dd, J = 11.3, 6.2 Hz, 2 H), 2.09 (m, 2 H). ¹³C NMR (101 MHz, CDCl₃): δ = 192.45, 167.20, 163.97, 161.53, 154.85, 144.66, 133.23, 129.73, 119.34, 116.35, 116.01, 114.62, 77.07, 67.47, 55.58, 54.43, 50.87, 50.22, 46.72, 44.32, 36.45, 23.70, 22.21. HRMS-ESI: m/z [M + H]⁺ calcd for C₂₇H₂₈O₃N₄F: 475.21482; found: 475.21400.

3-(4-Fluorophenyl)-2-[4-(4-nitrophenyl)piperazine-1-carbonyl]-4-oxo-2,3,4,5,6,7-hexahydro-1H-indole-2-carbonitrile (5p)

Yield: 176 mg (72%); yellow solid; mp 242–244 °C. ¹H NMR (400 MHz, CDCl₃): δ = 8.23–8.11 (m, 2 H), 7.33–7.28 (m, 2 H), 7.16–7.06 (m, 2 H), 6.93–6.83 (m, 2 H), 5.42 (s, 1 H), 4.45 (s, 1 H), 4.14 (d, J = 10.5 Hz, 1 H), 3.92 (s, 1 H), 3.70 (d, J = 44.9 Hz, 4 H), 3.49–3.31 (m, 2 H), 2.66–2.57 (m, 2 H), 2.32–2.24 (m, 2 H), 2.09 (m, 2 H). ¹³C NMR (101 MHz, CDCl₃): δ = 189.94, 167.29, 162.04, 159.41, 152.98, 136.82, 132.53, 128.71, 124.29, 114.62, 114.19, 113.98, 111.69, 110.54, 66.89, 52.00, 44.51, 43.83, 41.95, 34.84, 21.86, 20.67. HRMS-ESI: m/z [M + H]⁺ calcd for C₂₆H₂₅O₄N₅F: 490.18927; found: 490.18851.

3-(4-Fluorophenyl)-4-oxo-2-[4-(pyridin-2-yl)piperazine-1-carbonyl]-2,3,4,5,6,7-hexahydro-1H-indole-2-carbonitrile (5q)

Yield: 162 mg (73%); off-white solid; mp 178–180 °C. ¹H NMR (500 MHz, CDCl₃): δ = 8.23–8.19 (m, 1 H), 7.57–7.50 (m, 1 H), 7.33–7.28 (m, 2 H), 7.09 (t, J = 8.5 Hz, 2 H), 6.75–6.65 (m, 2 H), 5.69 (s, 1 H), 4.47 (s, 1 H), 4.12–3.39 (m, 8 H), 2.83–2.47 (m, 2 H), 2.37–2.18 (m, 2 H), 2.08 (dd, J = 12.4, 6.2 Hz, 2 H). ¹³C NMR (101 MHz, CDCl₃): δ = 192.44, 167.13, 164.05, 151.51, 135.17, 133.12, 130.30, 129.80, 120.90, 116.89, 116.40, 115.84, 114.80, 67.47, 54.42, 48.88, 48.30, 46.31, 43.99, 36.44, 23.70, 22.21. HRMS-ESI: m/z [M + H]⁺ calcd for C₂₅H₂₅O₂N₅F: 446.21482; found: 446.21400.

2-(4-Ethylpiperazine-1-carbonyl)-3-(4-fluorophenyl)-4-oxo-2,3,4,5,6,7-hexahydro-1H-indole-2-carbonitrile (5r)

Yield: 129 mg (65%); white solid; mp 142–144 °C. ¹H NMR (500 MHz, CDCl₃): δ = 7.29–7.25 (m, 1 H), 7.10–7.05 (m, 1 H), 5.54 (d, J = 3.8 Hz, 1 H), 4.42 (s, 1 H), 3.87 (d, J = 38.8 Hz, 1 H), 3.74–3.52 (m, 3 H), 2.69–2.55 (m, 4 H), 2.53–2.40 (m, 4 H), 2.29–2.24 (m, 2 H), 2.11–2.05 (m, 2 H), 1.11 (t, J = 7.2 Hz, 3 H). ¹³C NMR (101 MHz, CDCl₃): δ = 192.42, 167.21, 163.76, 161.50, 133.28, 129.79,

116.29, 116.07, 115.90, 67.42, 54.36, 52.09, 51.51, 46.63, 44.21, 36.45, 23.69, 22.20, 11.93. HRMS-ESI: m/z [M + H]⁺ calcd for C₂₂H₂₅N₄O₂F: 397.203979; found: 397.20519.

3-(4-Methoxyphenyl)-4-oxo-2-(piperidine-1-carbonyl)-2,3,4,5,6,7-hexahydro-1H-indole-2-carbonitrile (5s)

Yield: 117 mg (62%); white solid; mp 132–134 °C. ¹H NMR (500 MHz, CDCl₃): δ = 7.24–7.19 (m, 2 H), 6.93–6.87 (m, 2 H), 5.47 (s, 1 H), 4.38 (s, 1 H), 3.79 (s, 3 H), 3.78–3.71 (m, 1 H), 3.63 (d, J = 14.8 Hz, 1 H), 3.58–3.44 (m, 2 H), 2.59 (t, J = 8.0 Hz, 2 H), 2.28–2.23 (m, 2 H), 2.07 (dt, J = 11.9, 5.9 Hz, 2 H), 1.73 (s, 6 H). ¹³C NMR (126 MHz, CDCl₃): δ = 192.51, 167.05, 163.88, 159.63, 129.60, 129.10, 116.15, 114.47, 77.06, 67.76, 55.23, 54.39, 47.61, 45.44, 36.51, 25.38, 25.14, 24.11, 23.73, 22.25. HRMS-ESI: m/z [M + H]⁺ calcd for C₂₂H₂₆N₃O₃: 380.19720; found: 380.19687.

4-Oxo-2-(piperidine-1-carbonyl)-3-(3,4,5-trimethoxyphenyl)-2,3,4,5,6,7-hexahydro-1H-indole-2-carbonitrile (5t)

Yield: 128 mg (58%); white solid; mp 198–200 °C. ¹H NMR (400 MHz, CDCl₃): δ = 6.53 (s, 2 H), 5.49 (s, 1 H), 4.34 (s, 1 H), 3.85 (d, J = 6.1 Hz, 9 H), 3.71–3.52 (m, 4 H), 2.66–2.54 (m, 2 H), 2.33–2.17 (m, 2 H), 2.12–2.06 (m, 2 H), 1.82–1.61 (m, 6 H). ¹³C NMR (101 MHz, CDCl₃): δ = 192.47, 167.11, 163.82, 153.53, 138.34, 133.09, 115.96, 105.39, 67.63, 60.86, 56.27, 55.40, 47.59, 45.42, 36.54, 25.35, 24.09, 23.76, 22.30. HRMS-ESI: m/z [M + H]⁺ calcd for C₂₂H₂₆N₃O₅: 440.218000; found: 440.21798.

4-Oxo-3-phenyl-2-(piperidine-1-carbonyl)-2,3,4,5,6,7-hexahydro-1H-indole-2-carbonitrile (5u)

Yield: 115 mg (66%); white solid; mp 204–206 °C. ¹H NMR (500 MHz, CDCl₃): δ = 7.38 (ddd, J = 7.5, 4.4, 1.3 Hz, 2 H), 7.34–7.29 (m, 3 H), 5.44 (s, 1 H), 4.41 (s, 1 H), 3.80–3.71 (m, 1 H), 3.67–3.44 (m, 3 H), 2.60 (dd, J = 11.5, 5.7 Hz, 2 H), 2.29–2.23 (m, 2 H), 2.12–2.05 (m, 2 H), 1.62 (dt, J = 16.2, 7.7 Hz, 6 H). ¹³C NMR (101 MHz, CDCl₃): δ = 192.40, 167.30, 163.77, 137.55, 129.07, 128.63, 127.94, 116.01, 77.06, 67.61, 54.93, 47.59, 45.45, 36.47, 25.36, 25.11, 24.07, 23.69, 22.21. HRMS-ESI: m/z [M + H]⁺ calcd for C₂₁H₂₄N₃O₂: 350.18630; found: 350.18632.

4-Oxo-2-(piperidine-1-carbonyl)-3-(o-tolyl)-2,3,4,5,6,7-hexahydro-1H-indole-2-carbonitrile (5v)

Yield: 116 mg (64%); white solid; mp 162–164 °C. ¹H NMR (500 MHz, CDCl₃): δ = 7.43 (d, J = 7.5 Hz, 1 H), 7.24–7.14 (m, 3 H), 5.48 (s, 1 H), 4.67 (s, 1 H), 3.98 (s, 1 H), 3.44 (dd, J = 105.7, 41.0 Hz, 3 H), 2.58 (s, 3 H), 2.57–2.53 (m, 2 H), 2.21 (dt, J = 8.2, 4.6 Hz, 2 H), 2.03 (dd, J = 11.4, 5.2 Hz, 2 H), 1.68 (d, J = 21.4 Hz, 6 H). ¹³C NMR (101 MHz, CDCl₃): δ = 192.31, 166.43, 164.05, 136.46, 135.59, 130.62, 128.19, 127.93, 126.78, 118.52, 116.44, 67.69, 50.22, 47.54, 45.61, 36.48, 25.31, 24.12, 23.74, 22.05, 19.72. HRMS-ESI: m/z [M + H]⁺ calcd for C₂₂H₂₆N₃O₂: 364.20351; found: 364.20195.

3-(4-Nitrophenyl)-4-oxo-2-(piperidine-1-carbonyl)-2,3,4,5,6,7-hexahydro-1H-indole-2-carbonitrile (5w)

Yield: 106 mg (54%); white solid; mp 170–172 °C. ¹H NMR (400 MHz, CDCl₃): δ = 8.29–8.22 (m, 2 H), 7.52–7.47 (m, 2 H), 5.57 (s, 1 H), 4.51 (s, 1 H), 3.82 (d, J = 13.0 Hz, 1 H), 3.51 (dt, J = 18.0, 12.1 Hz, 3 H), 2.63 (dd, J = 10.2, 5.5 Hz, 2 H), 2.27 (dd, J = 14.7, 6.9 Hz, 2 H), 2.10 (dd, J = 12.3, 6.1 Hz, 2 H), 1.62 (s, 6 H). ¹³C NMR (126 MHz, CDCl₃): δ = 192.32, 167.84, 163.03, 147.97, 144.88, 129.16, 124.42, 115.61, 115.53, 66.87, 54.45, 47.63, 45.67, 36.34, 25.35, 25.15, 24.02, 23.73, 22.17. HRMS-ESI: m/z [M + H]⁺ calcd for C₂₁H₂₂N₄O₄: 394.16420; found: 394.16340.

3-(4-Fluorophenyl)-6,6-dimethyl-4-oxo-2-(piperidine-1-carbonyl)-2,3,4,5,6,7-hexahydro-1H-indole-2-carbonitrile (5x)

Yield: 142 mg (72%); white solid; mp 160–162 °C. ¹H NMR (500 MHz, CDCl₃): δ = 7.29 (ddd, J = 7.0, 4.5, 2.0 Hz, 2 H), 7.10–7.05

(m, 2 H), 5.35 (s, 1 H), 4.40 (s, 1 H), 3.79 (d, $J = 13.2$ Hz, 1 H), 3.61 (d, $J = 13.4$ Hz, 1 H), 3.47 (dd, $J = 23.6, 14.8$ Hz, 2 H), 2.44 (d, $J = 5.8$ Hz, 2 H), 2.14 (dd, $J = 40.0, 16.4$ Hz, 2 H), 1.73 (d, $J = 4.7$ Hz, 6 H), 1.10 (d, $J = 2.3$ Hz, 6 H). ^{13}C NMR (101 MHz, CDCl_3): $\delta = 191.83, 166.33, 163.92, 163.61, 161.46, 133.53, 129.64, 116.26, 116.05, 115.95, 114.47, 67.67, 54.11, 50.57, 47.60, 45.51, 37.42, 34.65, 29.07, 28.28, 25.37, 25.10, 24.06$. HRMS-ESI: m/z [M + H]⁺ calcd for $\text{C}_{23}\text{H}_{27}\text{O}_2\text{N}_3\text{F}$: 396.20818; found: 396.20904.

3-(4-Fluorophenyl)-6,6-dimethyl-4-oxo-2-[4-(pyridin-2-yl)piperazine-1-carbonyl]-2,3,4,5,6,7-hexahydro-1H-indole-2-carbonitrile (5y)

Yield: 172 mg (73%); white solid; mp 164–166 °C. ^1H NMR (500 MHz, CDCl_3): $\delta = 8.23\text{--}8.19$ (m, 1 H), 7.53 (ddd, $J = 8.9, 7.2, 2.0$ Hz, 1 H), 7.34–7.29 (m, 2 H), 7.13–7.07 (m, 2 H), 6.74–6.70 (m, 1 H), 6.68 (d, $J = 8.6$ Hz, 1 H), 5.41 (s, 1 H), 4.45 (s, 1 H), 3.98 (d, $J = 12.9$ Hz, 1 H), 3.84 (t, $J = 9.0$ Hz, 3 H), 3.75–3.55 (m, 3 H), 3.50–3.40 (m, 1 H), 2.46 (s, 2 H), 2.18–2.09 (m, 2 H), 1.11 (d, $J = 4.3$ Hz, 6 H). ^{13}C NMR (101 MHz, CDCl_3): $\delta = 191.79, 165.99, 164.21, 158.72, 148.08, 137.91, 133.24, 129.79, 116.42, 116.21, 115.92, 114.57, 107.46, 67.68, 54.35, 50.62, 46.23, 44.98, 44.50, 43.99, 37.46, 34.70, 28.97, 28.38$. HRMS-ESI: m/z [M + H]⁺ calcd for $\text{C}_{27}\text{H}_{29}\text{O}_2\text{N}_5\text{F}$: 474.23108; found: 474.22998.

3-(4-Fluorophenyl)-4-oxo-6-phenyl-2-[4-(pyridin-2-yl)piperazine-1-carbonyl]-2,3,4,5,6,7-hexahydro-1H-indole-2-carbonitrile (5z)

Yield: 164 mg (63%); white solid; mp 210–212 °C. ^1H NMR (500 MHz, CDCl_3): $\delta = 8.29\text{--}8.13$ (m, 1 H), 7.60–7.48 (m, 1 H), 7.39–7.31 (m, 2 H), 7.25 (m, 5 H), 7.10 (m, 2 H), 6.72 (dd, $J = 7.1, 4.9$ Hz, 1 H), 6.69 (d, $J = 8.5$ Hz, 1 H), 5.58 (d, $J = 15.4$ Hz, 1 H), 4.50 (d, $J = 7.9$ Hz, 1 H), 3.98 (d, $J = 12.5$ Hz, 1 H), 3.80 (t, $J = 18.3$ Hz, 3 H), 3.76–3.56 (m, 3 H), 3.53–3.40 (m, 2 H), 2.90–2.79 (m, 2 H), 2.57–2.54 (m, 2 H). ^{13}C NMR (101 MHz, CDCl_3): $\delta = 191.03, 164.08, 158.68, 148.03, 142.26, 137.85, 129.84, 128.81, 127.18, 126.83, 126.72, 116.29, 116.08, 115.76, 114.52, 107.44, 77.06, 67.66, 54.36, 46.20, 44.94, 44.47, 43.93, 43.51, 41.07, 40.22, 31.28, 30.89$. HRMS-ESI: m/z [M + H]⁺ calcd for $\text{C}_{31}\text{H}_{29}\text{O}_2\text{N}_5\text{F}$: 522.23097; found: 522.22998.

3-(4-Chlorophenyl)-2-cyano-*N,N*-diethyl-4-oxo-2,3,4,5,6,7-hexahydro-1H-indole-2-carboxamide (5a')

Yield: 190 mg (52%); colorless liquid. ^1H NMR (400 MHz, CDCl_3): $\delta = 7.34\text{--}7.30$ (m, 2 H), 7.27 (d, $J = 2.6$ Hz, 1 H), 7.23–7.19 (m, 1 H), 5.69 (s, 1 H), 4.38–4.35 (m, 1 H), 3.54 (dt, $J = 13.5, 6.8$ Hz, 1 H), 3.50–3.37 (m, 3 H), 2.59 (dd, $J = 12.4, 6.2$ Hz, 2 H), 2.28 (dd, $J = 6.9, 4.7$ Hz, 2 H), 2.08 (dt, $J = 12.6, 6.2$ Hz, 2 H), 1.33 (t, $J = 6.9$ Hz, 3 H), 1.19 (t, $J = 7.1$ Hz, 3 H). ^{13}C NMR (126 MHz, CDCl_3): $\delta = 192.33, 167.80, 164.26, 139.64, 134.89, 130.33, 128.91, 128.06, 126.25, 115.81, 115.20, 67.54, 54.56, 42.27, 41.64, 36.41, 23.65,$

22.17, 12.82, 12.40. HRMS-ESI: m/z [M + H]⁺ calcd for $\text{C}_{20}\text{H}_{23}\text{N}_3\text{O}_2\text{Cl}$: 372.14882; found: 372.14956.

1-[4-(4-Fluorophenyl)piperazin-1-yl]ethan-1-one (7)

^1H NMR (400 MHz, CDCl_3): $\delta = 7.02\text{--}6.94$ (m, 2 H), 6.92–6.83 (m, 2 H), 3.81–3.73 (m, 2 H), 3.67–3.57 (m, 2 H), 3.13–2.97 (m, 4 H), 2.14 (s, 3 H). ^{13}C NMR (101 MHz, CDCl_3): $\delta = 169.05, 158.73, 156.35, 147.52, 118.59, 115.85, 115.53, 50.68, 50.32, 46.22, 41.35, 29.64, 21.30$. HRMS-ESI: m/z [M + H]⁺ calcd for $\text{C}_{12}\text{H}_{16}\text{ON}_2\text{F}$: 223.12412; found: 223.12470.

Crystal Data for 5a

$\text{C}_{24}\text{H}_{22}\text{N}_3\text{O}_2\text{Cl}$, $M = 401.88$, monoclinic, space group $P2_1/n$ (No.14), $a = 14.005(12)\text{Å}$, $b = 14.440(12)\text{Å}$, $c = 10.998(9)\text{Å}$, $\alpha = 90^\circ$, $\beta = 112.583(18)^\circ$, $\gamma = 90^\circ$, $V = 2054(3)\text{Å}^3$, $Z = 4$, $D_c = 1.300\text{ g/cm}^3$, $F_{000} = 848$, Bruker D8 QUEST PHOTON-100, Mo $K\alpha$ radiation, $\lambda = 0.71073\text{ Å}$, $T = 293(2)\text{ K}$, $2\theta_{\text{max}} = 50^\circ$, $\mu = 0.212\text{ mm}^{-1}$, 13591 reflections collected, 3595 unique ($R_{\text{int}} = 0.0667$), 265 parameters, $R1 = 0.0618$, $wR2 = 0.1542$, R indices based on 2598 reflections with $I > 2\sigma(I)$ (refinement on F^2), final $\text{Goof} = 1.025$, largest difference hole and peak = -0.390 and 0.570 e Å^{-3} (Figure 1). CCDC 2065225 contains the supplementary crystallographic data for this paper. The data can be obtained free of charge from The Cambridge Crystallographic Data Centre via www.ccdc.cam.ac.uk/structures

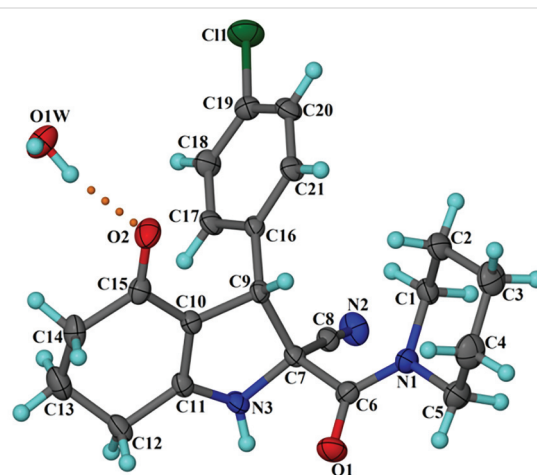


Figure 1 ORTEP diagram of compound 5a with atom numbering. The compound crystallized as monohydrate. Displacement ellipsoids are drawn at the 35% probability level and H atoms are shown as small spheres of arbitrary radius. Hydrogen bonds are shown in dotted lines.