Imido-substituted triazines as dehydrative condensing reagents for the chemoselective formation of amides in the presence of free hydroxy groups†

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In this paper, we discuss the synthesis of imido-substituted chlorotriazines and demonstrate their use in dehydrative condensation reactions. Chemoselective amide-forming reactions of amino alcohols using succinimido-substituted chlorotriazine (2A) proceeded smoothly. Occasionally, nonselectivity was problematic during the synthesis of hydroxy-substituted amides. Moreover, it was noteworthy that this method was applicable to hydroxy-substituted carboxylic acids that could have formed a lactone or an ester during the carboxylic acid activation step. The imido-substituted chlorotriazine (2A) was superior to the amido-substituted chlorotriazine and 2-chloro-4,6-dimethoxy-1,3,5-triazine (CDMT) in terms of reaction rates and yields.

Introduction

Amides are important functional groups that can be found in many materials, pharmaceuticals, and natural products. Therefore, they are routinely synthesized from carboxylic acids and amines in both the laboratory and industry.1 Chemoselective amide-forming reactions of amino alcohols have long been investigated2–4 because many important compounds contain this combination of functional groups, including anti-inflammatory drugs such as acetaminophen,2 cosmetics,3 and surfactants.4 However, nonselective N- and O-acylation of amino alcohols is often problematic during amide synthesis.2–4,7 In addition, the activation of a hydroxy-substituted carboxylic acid is difficult using conventional methods because lactone or ester formation is possible during carboxylic acid activation.

We have developed a class of triazine-based dehydrative condensing reagents [(4,6-dimethoxy-1,3,5-triazin-2-yl)trialkylammonium salts, DMT-Ams], including 4-(4,6-dimethoxy-1,3,5-triazin-2-yl)-4-methylmorpholinium chloride (DMT-MM), which can be employed in aqueous or alcoholic solvents (Scheme 1. See the ESI† for the detailed reaction mechanism).8,9 These reagents were prepared from 2-chloro-4,6-dimethoxy-1,3,5-triazine (CDMT) and N-methylmorpholine (NMM) or tertiary amines (tert-amines).9 For the preparation of functional DMT-Ams that selectively activate carboxylic acids, various tert-amines were studied.10,11 However, in most cases, the substituents on the triazine ring were limited to dimethoxy groups. Considering that substituents on the triazine would be important for the development of a diverse family of dehydrative condensing reagents, we studied chlorotriazines and their ammonium salts. As part of this project, we have recently reported that electron-withdrawing amido-substituted triazines (1 and 3) showed a higher reactivity for dehydrative condensing reactions.12 According to the Hammett substituent constants,13 imido-substituents (σ_m = 0.34 for the succinimido group) are more electron-withdrawing than methoxy- (σ_m = 0.12) and...
amido-substituents ($\sigma_m = 0.21$ for the acetamido group); therefore, a higher reactivity for condensation was expected. In this study, triazines with even more electron-withdrawing imido-substituents (2 and 4) were investigated, including the chemoselective acylation of amino alcohols and activation of hydroxy-substituted carboxylic acids.

Results and discussion

Rapidly available imides, such as succinimide, phthalimide, maleimide, and glutarimide, as substituents for triazine, were studied because condensing reagents should be inexpensive, especially if they are to be used in large-scale syntheses (2A–2D in Fig. 1). Moreover, instead of cyclic imides, linear diacetamide was also considered (2E in Fig. 1). For simplification, only one methoxy group on DCMT was replaced with an imido-substituent (Fig. 1).

It was difficult to determine the general reaction conditions for the synthesis of the above-mentioned imido-substituted chlorotriazines because the different imides possessed unequal reactivities (Scheme 2). The succinimido-substituted compound 2A was obtained from the reaction of 2,4-dichloro-6-methoxy-1,3,5-triazine (DCMT) with the succinimide anion in tetrahydrofuran (THF) at 55 °C (55% isolated yield). 2B was prepared in a 58% isolated yield using the commercially available phthalimide potassium salt and DCMT in the presence of 18-crown-6 at ambient temperature. However, 2C could not be obtained under similar reaction conditions using DCMT in the presence of various bases. Compared to the five-membered ring succinimide, the nucleophilicity of the six-membered glutarimide anion toward DCMT was unexpectedly low. Despite the decreased delocalization of the negative charge for the glutarimide anion as expected from their $pK_a$ values (11.4 for glutarimide and 9.6 for succinimide), the yield of 2D was low.\textsuperscript{14} Shielding of the nitrogen lone pairs of the imide anion by the two adjacent carbonyl oxygen groups, as expected from X-ray crystal structure analysis of glutarimide\textsuperscript{15} and succinimide,\textsuperscript{16} may explain the reduced reactivity. Glutarimide had a smaller $\theta$–$\mathrm{N}$ bond angle (119.60° for glutarimide and 123.86° for succinimide) and larger C–N–C bond angle (126.90° for glutarimide and 112.62° for succinimide). 2D was successfully crystallized and the structure was confirmed by X-ray single crystal structure analysis (see the ESI†). In the case of the linear diaacetamide, 2E was difficult to prepare. Despite the consumption of DCMT, only side products were detected.

Next, we examined the preparation of the imido-substituted condensing reagents (4). Although the desired 4 would be formed using NMM and N-methylpiperidine (NMP) as the tert-amines, the purities of 4 were low because demethylation (by the chloride anion) and hydrolysis occurred during the isolation steps. To prevent demethylation, the synthesis of 4 was attempted in the presence of nonnucleophilic perchlorate anions; however, hydrolysis was still problematic. It was assumed that the strong electron-withdrawing effect of the imido group made isolation difficult.

For convenient evaluation of the reactivity of these prepared chlorotriazines (2A, 2B, and 2D), the corresponding DMT-Ams (10) were generated in situ and used for coupling of carboxylic acid 5a and amine 6a (Scheme 3 and Table 1).\textsuperscript{9} Chlorotriazine 2A was added to a mixture of 5a, 6a, and NMM at ambient temperature in MeOH (Table 1, entry 1).\textsuperscript{17} Amide 7a was only obtained in a 69% yield. The formation of side product 8a (3%) and methyl ester 9a (11%) was also observed. The yield of 9a using 2A was larger than that using CDMT (2% for CDMT), indicating that the electrophilicity of triazinyl ester 11 toward methanol was increased because of the succinimido group on the triazine ring. In THF, the increased yield of the side product 8a (35% for 2A) was observed, in contrast to CDMT (18% for CDMT), implying a similar electron-withdrawing effect of the
succinimido group. To prevent the alcoholysis of 11, the reaction was conducted in 1PrOH and the yield of 7a was improved to 84% (Table 1, entry 1). Due to the fact that similar results were obtained for 2B and 2D (Table 1, entries 2 and 3), the influence from the kind of imido-structures was thought to be small for the condensing reactions. Moreover, the phthalimido-substituted chlorotriazine (2B) was not very soluble in THF and alcohol; therefore, only succinimido-substituted chlorotriazine 2A and glutarimido-substituted chlorotriazine 2D were selected for further studies.

The formation of the side product 8 was assumed to result from chlorotriazine 2 being more electrophilic than 10 or 11. Thus, the addition of amine 6 after the formation of 11 should prevent the formation of 8 (Table 2). Accordingly, amine 6 was added to the preformed mixture of chlorotriazine, NMM, and carboxylic acid 5 in THF. In this solvent, ester 9 would never be formed. In the amide-forming reactions of 2-phenylpropionic acid (5a) with 2-phenethylamine (6a), aniline (6b), or 4-(ethoxycarbonyl)aniline (6c) (the nucleophilicity of these amines decreases in this order), imido-substituted 2A and 2D afforded better yields than CDMT (Table 2, entries 1–9). From the viewpoints of atom economy, starting material costs, and synthetic yields, 2A was selected for further studies. In the case of sterically hindered isobutyric acid (5b, entries 10 and 11), 2A also afforded amide 7d in a better yield than that using CDMT. When the more sterically hindered pivalic acid (5e) was used (entries 12–17), different reactivities between 2A and CDMT were observed for aniline 6b (entry 16 versus 17). Although it was thought that the benzoic acid derivative (5f, with electron-withdrawing nitro groups) might form intermediate 11 slowly, 2A afforded product 7h in an excellent yield compared to CDMT (Table 2, entry 12 versus 19). This method is also effective for formation of a peptide bond, i.e., condensation of Boc-Leu-OMe (5e) and H-Phe-OMe (6e) afforded the corresponding dipeptide (7j) in 94% NMR yield as a single diastereomer (Table 2, entry 20). As indicated in entry 18 of Table 2, purification by only extraction and recrystallization without column chromatography was also possible.

From these results, it is unambiguous that the succinimido-substituted chlorotriazine 2A was more reactive than CDMT, and less sterically hindered alcoholic solvents (especially methanol) are unsuitable for carrying out the condensing reactions because of alcoholysis of 11. However, we anticipated that the competing alcoholysis against aminolysis would be reduced when lower concentrations of alcohols were used. Namely, chemoselective amide forming reactions could be effective even using carboxylic acids 5 and amines 6 having free hydroxy groups. Thus, we tested 6-hydroxyhexanoic acid (5f) as it would be susceptible to an intramolecular lactone-formation or an intermolecular ester-formation during the carboxylic acid activation step (Table 3, entries 1 and 2). The desired amide 7j was obtained in a good yield compared to CDMT under the reaction conditions used in Table 2. When amino alcohols such as benzyl(2-hydroxyethyl)amine (6f, Table 3, entries 3 and 4) and N-(2-hydroxyethyl)aniline (6g, Table 3, entries 5 and 6) were employed, better yields of amide products (7k and 7l) were
observed, even in the presence of the competing free hydroxy groups. Although the chemoselective acylation of 4-amino-phenol (6h) using EDC and HOBt in the presence of a surfactant was insufficient, 2A afforded the desired product 7m chemoselectively. Even though Boc-Tyr-OH having the phenolic hydroxy group was used as the starting material, the desired dipeptide (7n) was obtained in 80% NMR yield and 71% isolated yield (Table 3, entry 9).

To clarify the reactivity of 2A compared with the amido-substituted chlorotriazine 1A and CDMT, the formation of 7l under similar conditions used in Tables 2 and 3 was investigated kinetically with 1H NMR spectroscopy (Fig. 2). The reaction rate with 2A was faster than that with 1A or CDMT.

From the study of the DMT-Ams, we previously elucidated that the reactivity of tert-amines toward CDMT was strongly influenced by their steric hinderance; that is, a β-alkyl group toward the nitrogen lone pair in a gauche relationship remarkably hampers the nucleophilic attack on CDMT. Because of this effect, N-ethylmorpholine, N-ethylpiperidine, and triethylamine are unable to react with CDMT to afford DMT-Ams at ambient temperature. Contrastingly, the imido-
substituted chlorotriazine 2A reacted with these tert-amines, resulting in the formation of 7a (Table 4). The strong electron-withdrawing ability of the succinimido group was thought to be responsible for overcoming this difficulty, thus enabling the use of various tert-amines for condensing reagents.10,11

**Conclusion**

We have developed imido-substituted chlorotriazines (2) and demonstrated their use in efficient dehydrative condensing reactions between carboxylic acids and amines. Chemoselective amide-forming reactions using amino alcohols with

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**Table 3** Amide-forming reactions in the presence of free hydroxy groups

| Entry | Chlorotriazine | Carboxylic acid 5 | Amine 6 | Amide 7 | Time | Yieldb (%) |
|-------|----------------|-------------------|---------|---------|------|------------|
| 1     | 2A             | 5f                | 6c      |         | 4 h  | 81 (75)    |
| 2     | CDMT           |                   |         |         | 4 h  | 30         |
| 3     | 2A             | 5a                | 6f      |         | 15 min | 88 (84) |
| 4     | CDMT           |                   | 6f      |         | 15 min | 71         |
| 5     | 2A             | 5a                | 6g      |         | 6 h  | 97 (85)    |
| 6     | CDMT           |                   | 6g      |         | 6 h  | 73         |
| 7b    | 2A             | 5a                | 5g      | H-Gly-O'Bu | 1 h | 90 (73)    |
| 8b    | CDMT           |                   | 5a      | H-Gly-O'Bu | 15 min | 54        |
| 9c    | 2A             |                   | 5g      | Boc-Tyr-Gly-O'Bu | 1 h | 80 (71)    |

**Table 4** Amide-forming reactions using sterically hindered tert-amines that were incapable of reacting with CDMT

| Entry | tert-Amine | Time (min) | Yieldb (%) |
|-------|------------|------------|------------|
| 1     |            | 30         | 78         |
| 2     |            | 30         | 77         |
| 3     |            | 60         | 62         |
| 4     | Et3N       | 60         | 63         |

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*a* NMR yields. Isolated yields are given in the parentheses. *b* 3-Phenylpropionic acid (1.1 eq.), 2A or CDMT (1.0 eq.), NMM (1.05 eq.), and 4-aminophenol (1.0 eq.) were used. *c* Boc-Tyr-OH (1.0 eq.), 2A (1.0 eq.), NMM (1.2 eq.), H-Gly-O'Bu-HCl (2.0 eq.), and EtN'Pr₂ (2.0 eq.) were used.

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**Fig. 2** Kinetic study of amide 7l formation using 2A, 1A, and CDMT.
succinimido-substituted 2A were successfully achieved. In addition, we demonstrated that 2A was an effective reagent for the chemoselective condensing reaction of hydroxy-substituted carboxylic acids that could form a lactone or an ester during the activation step. In terms of reaction rates and amide yields, 2A was superior to the amido-substituted chlorotriazine 1A and CDMT. Sterically hindered tert-amines that would never react with CDMT at ambient temperature, such as N-ethylmorpholine, N-ethylpiperidine, and triethylamine, were successfully employed in the formation of DMT-Ams with 2A. These results will contribute to the development of novel functionalized triazine-based condensing reagents and tert-amine-catalyzed amide-forming reactions.

**Experimental**

**General methods**

Nuclear magnetic resonance ([1H NMR (400 or 600 MHz) and [13C NMR (100 or 150 MHz)]) spectroscopy was carried out on JEOL JNM-ECS400 and JEOL JNM-ECA600 spectrometers. Chemical shifts for [1H NMR were reported as δ values relative to tetramethylsilane (TMS). The coupling constants for [1H NMR were reported in hertz (Hz). The following abbreviations were used for reporting the spin multiplicity: s = singlet, d = doublet, t = triplet, q = quartet, quin = quintet, m = multiplet, and br = broad. Chemical shifts for [13C NMR were reported as δ values relative to the center line of the deuterochloroform triplet at 77.16 ppm. Mass spectra were measured on a JEOL JMS-T100TD spectrometer (DART- and ESI-MS). Analytical thin-layer chromatography (TLC) was performed on Merck precoated analytical plates. 0.25 mm thick, silica gel 60 F254. Flash chromatography separation was performed using Kanto Chemical silica gel 60 N (spherical, neutral, 40-100 mesh) unless otherwise noted. All Reagents were of commercial grade and were used without further purification unless otherwise noted. Anhydrous THF and toluene were purchased from commercial sources. All reactions that are sensitive to oxygen or moisture were conducted under a N2 atmosphere. The [1H NMR spectral data for 2-methoxy-4-phénylamino-6-methyl-1,3,5-triazine (8A) and 2-glutarimido-4-methoxy-6-phénylamino-1,3,5-triazine (8D) were consistent with those of authentic samples prepared from 2A and 2D (see the ESI† for synthetic details). [1H NMR spectral data for methyl 3-phenylpropanoate (9a) and 2-propyl 3-phenylpropanoate (9b) were consistent with the referenced chemical shift values. [1H NMR spectral data of the isolated amide products 7a, 7b, 7c, 7d, 7e, 7f, 7g, 7h, 7i, 7m, 7n and 7i (ref. 31) were identical to known literature values (see the ESI†).

**Experimental procedure and characterization data**

**2-Chloro-4-methoxy-6-succinimido-1,3,5-triazine (2A).** To a THF solution (70 mL) of sodium hydride (60%, 1.44 g, 36.0 mmol), succinimide (3.27 g, 33.0 mmol) was added at 0 °C. To this reaction mixture, a THF solution (0.3 mL) of cyanuric chloride (369 mg, 2.0 mmol) were sequentially added at 0 °C. The reaction mixture was quenched with 1 M KHSO4 and extracted with AcOEt. The organic phase was washed with brine, dried over anhydrous sodium sulfate, filtered and washed with AcOEt. After concentrating the filtrate under reduced pressure, the residue was purified by flash column chromatography (hexane/AcOEt = 2 : 1) to afford 2B (1.68 g, 58% yield) as a white solid. Mp 163-165 °C; [1H NMR (400 MHz, CDCl3): δ 8.02 (dd, J = 5.5, 3.0 Hz, 2H), 7.87 (dd, J = 5.5, 3.0 Hz, 2H), 4.17 (s, 3H); [13C NMR (100 MHz, CDCl3): δ 173.19, 172.40, 164.94, 162.76, 135.46, 131.29, 124.67, 56.74; IR (KBr): 3095, 3064, 2958, 1797, 1739, 1558, 1419, 1353, 1271, 1082, 935 cm⁻¹; HRMS (DART) m/z: [M + H]⁺ calcd for C12H11ClN4O3 260.0281; found: 260.0284; anal. calcd for C12H10ClN4O3: C, 39.43; H, 2.90; N, 22.89. Found: C, 40.11; H, 2.50; N, 19.23.

**To a toluene solution (100 mL) of 2,4-dichloro-6-methoxy-1,3,5-triazine (2.70 g, 15.0 mmol), potassium phthalimide (2.78 g, 15.0 mmol) and 18-crown-6 (0.39 g, 1.5 mmol) were added at ambient temperature. After stirring for 12 h, the reaction mixture was filtered and washed with AcOEt. After concentrating the filtrate under reduced pressure, the residue was purified by flash column chromatography (hexane/AcOEt = 2 : 1) to afford 2B (1.68 g, 58% yield) as a white solid. Mp 163-165 °C; [1H NMR (400 MHz, CDCl3): δ 8.02 (dd, J = 5.5, 3.0 Hz, 2H), 7.87 (dd, J = 5.5, 3.0 Hz, 2H), 4.17 (s, 3H); [13C NMR (100 MHz, CDCl3): δ 173.19, 172.40, 164.94, 162.76, 135.46, 131.29, 124.67, 56.74; IR (KBr): 3095, 3064, 2958, 1797, 1739, 1558, 1419, 1353, 1271, 1082, 935 cm⁻¹; HRMS (DART) m/z: [M + H]⁺ calcd for C12H11ClN4O3 260.0281; found: 260.0284; anal. calcd for C12H10ClN4O3: C, 39.43; H, 2.90; N, 22.89. Found: C, 40.11; H, 2.50; N, 19.23.

**2-Chloro-4-glutarimido-6-methoxy-1,3,5-triazine (2D).** To a THF solution (1 mL) of sodium hydride (88 mg, 2.2 mmol), glutarimide (272 mg, 2.4 mmol) in THF (5.5 mL) was added at ambient temperature. To this mixture, a THF solution (0.3 mL) of 2,4-dichloro-6-methoxy-1,3,5-triazine (2.70 g, 15.0 mmol) was sequentially added dropwise at −78 °C. After stirring for 7 h at ambient temperature, methanol (10 mL) and NaHCO3 (336 mg, 4.0 mmol) were added at 0 °C. After stirring for 13 h at ambient temperature, the reaction mixture was quenched with 1 M KHSO4 (2 mL) at 0 °C and extracted with CH2Cl2. The organic phase was washed with 1 M KHSO4 and brine, dried over anhydrous sodium sulfate, and concentrated under reduced pressure. The residue was purified by flash column chromatography (hexane/AcOEt = 9 : 1) to afford 2D (135 mg, 26% yield) as a white solid. Mp 125-127 °C; [1H NMR (400 MHz, CDCl3): δ 4.14 (s, 3H), 2.79 (t, J = 6.4 Hz, 4H), 2.14 (quin, J = 6.4 Hz, 2H); [13C NMR (100 MHz, CDCl3): δ 173.85, 173.13, 171.19, 167.12, 57.00, 32.35, 17.11; IR (KBr): 3016, 2970, 1743, 1697, 1560, 1539, 1498, 1408, 1385, 1282, 1248, 1144, 1016, 920, 760, 561, 442 cm⁻¹; HRMS (DART) m/z: [M + H]⁺ calcd for C13H12ClN4O3 275.0441; found: 275.0444; anal. calcd for C13H11ClN4O3: C, 42.12; H, 3.53; N, 21.83. Found: C, 42.05; H, 3.47; N, 21.83.

**General procedure for the amide-forming reactions (Tables 2, 3, and Fig. 2).** To a THF solution (2 mL) of carboxylic acid 5 (0.4 mmol) and NMM (53 µL, 0.48 mmol), chlorotriazine 2 or CDMT (0.44 mmol) was added. After stirring for 5 min at ambient temperature, amine 6 (0.44 mmol) was added to the
reaction mixture. The reaction was monitored for completion using TLC and quenched with 1 M KHSO₄ (4 mL). The mixture was extracted with CHCl₃ (10 mL × 3), and the combined organic layers were washed with 1 N HCl, sat. NaHCO₃, and brine. The organic layer was dried over Na₂SO₄, filtered, and concentrated under reduced pressure, and the residue was analyzed by quantitative NMR using coumarin or 1,3,5-trimesitylbenzene as an internal standard. After concentration under reduced pressure, the residue was purified by flash column chromatography with the appropriate eluent to afford amide 7.

**N-(4-Ethoxycarbonylphenyl)-3-phenylpropionamide (7c).**

The title compound (107 mg, 90% yield) was obtained by flash column chromatography (hexane/AcOEt = 8 : 2) as a white solid. Mp 136–137 °C; ¹H NMR (400 MHz, CDCl₃): δ 7.97 (d, J = 8.8 Hz, 2H), 7.52 (d, J = 8.8 Hz, 2H), 7.52 (br, 1H), 7.30–7.20 (m, 5H), 4.34 (q, J = 7.2 Hz, 2H), 3.04 (t, J = 7.6 Hz, 2H), 2.69 (t, J = 7.6 Hz, 2H), 1.37 (t, J = 7.2 Hz, 3H); ¹³C NMR (100 MHz, CDCl₃): δ 170.85, 166.32, 142.03, 140.49, 138.85, 130.85, 128.81, 128.47, 126.60, 125.97, 118.93, 61.03, 39.60, 31.50, 14.45, IR (KBr): 3315, 2935, 1454, 1408, 1273, 1192, 1074, 1059, 1028, 858, 772, 769, 752, 700, 652, 513 cm⁻¹; HRMS (ESI) [M + Na]⁺ calcd for C₁₅H₁₃N₃O₅: 316.0934; found: 316.0936; anal. calcd for C₁₅H₁₃N₃O₅: C, 57.14; H, 4.70; N, 13.33. Found: C, 56.92; H, 4.15; N, 13.31.

**N-(4-Ethoxycarbonylphenyl)-6-hydroxyhexanamide (7j).**

The title compound (80 mg, 75% yield) was obtained by flash column chromatography (hexane/AcOEt = 9 : 1 to CHCl₃, CHCl₃/AcOEt = 7 : 3) as a white solid. Mp 155–156 °C; ¹H NMR (400 MHz, CDCl₃): δ 9.14 (t, J = 2.0 Hz, 1H), 8.86 (d, J = 2.0 Hz, 2H), 7.36 (t, J = 7.6 Hz, 2H), 7.29–7.24 (m, 3H), 6.43 (br, 1H), 3.80 (td, J = 6.9, 6.9 Hz, 2H), 3.00 (t, J = 6.9 Hz, 2H); ¹³C NMR (100 MHz, CDCl₃): δ 162.91, 148.76, 138.24, 138.16, 129.09, 128.89, 127.22, 127.16, 121.19, 41.83, 35.50, IR (KBr): 3415, 3097, 2945, 1672, 1630, 1603, 1541, 1527, 1487, 1452, 1435, 1346, 1327, 1306, 1282 cm⁻¹; HRMS (DART) [M + H]⁺ calcd for C₁₅H₁₉N₂O₃: 292.1094; found: 292.1093; anal. calcd for C₁₅H₁₉N₂O₃: C, 61.14; H, 5.56; N, 13.33. Found: C, 61.16; H, 5.47; N, 13.28.

**3,5-Dinitro-N-(2-phenethyl)benzamide (7h).**

The title compound (119 mg, 94% yield) was obtained by flash column chromatography (hexane/AcOEt = 9 : 1 to CHCl₃, CHCl₃/AcOEt = 7 : 3) as a white solid. Mp 155–156 °C; ¹H NMR (400 MHz, CDCl₃): δ 9.14 (t, J = 2.0 Hz, 1H), 8.86 (d, J = 2.0 Hz, 2H), 7.36 (t, J = 7.6 Hz, 2H), 7.29–7.24 (m, 3H), 6.43 (br, 1H), 3.80 (td, J = 6.9, 6.9 Hz, 2H), 3.00 (t, J = 6.9 Hz, 2H); ¹³C NMR (100 MHz, CDCl₃): δ 162.91, 148.76, 138.24, 138.16, 129.09, 128.89, 127.22, 127.16, 121.19, 41.83, 35.50, IR (KBr): 3415, 3097, 2945, 1672, 1630, 1603, 1541, 1527, 1487, 1452, 1435, 1346, 1327, 1306, 1282 cm⁻¹; HRMS (DART) [M + H]⁺ calcd for C₁₅H₁₉N₂O₃: 292.1094; found: 292.1093; anal. calcd for C₁₅H₁₉N₂O₃: C, 61.14; H, 5.56; N, 13.33. Found: C, 61.16; H, 5.47; N, 13.28.
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