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A novel acid-catalyzed rearrangement of 2-substituted-3-(2-nitrophenyl)oxiranes for the synthesis of di- and mono-oxalamides†

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A novel one-pot synthetic approach to *N*¹-(2-carboxyaryl)-*N*²-(aryl or *H*)oxalamides from 3-(2-nitroaryl)oxirane-2-carboxamides via the classical Meinwald rearrangement and a new rearrangement sequence has been developed. The methodology is applicable to the synthesis of *N*-(2-carboxyphenyl)aryloxalmonoamides from (3-(2-nitrophenyl)oxiran-2-yl)(aryl)methanones. The method is operationally simple and high yielding, thus providing a new useful formula for both anthranilic acid derivatives and oxalamides.

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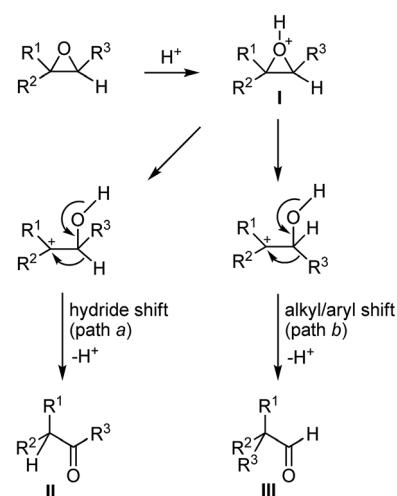
Introduction

Oxiranes are one of the most versatile classes of organic compounds available to the synthetic chemist.¹ They can be prepared by a wide variety of methods.² One of the most frequently used atom economical reactions of oxiranes is their rearrangement to carbonyl compounds, and a number of reagents including a variety of Lewis acids³ have been elaborated for this purpose. In principle, for trisubstituted oxiranes two types of rearrangements are possible depending on the migration pathways following the Lewis acid promoted C–O bond cleavage (Scheme 1). The rearrangement of **I** with hydride (path a) or the alkyl/aryl migration (path b) would lead to ketone **II** or aldehyde **III**, respectively.^{3a,4} The synthetic applications of oxiranes have been the subject of a number of reviews.^{1a,2,5}

The promise of increased chemo-, regio-, and stereoselectivity available *via* transition metal catalysis⁶ has led investigators to study the interactions of oxiranes with transition metal complexes, and a number of interesting and useful isomerization reactions have been reported. Notably, oxiranes activated by adjacent aryl, vinyl, silyl, or carbonyl substituents

are isomerized to carbonyl compounds or allylic alcohols by complexes of Rh,⁷ Pd,⁸ Mo,⁹ Sm,¹⁰ Fe¹¹ and In.¹²

Unlike all the considered reactions proceeding depending on the structure of oxirane and applied conditions of rearrangement on the path a or b, our strategy included the use of oxiranes, containing substituents with functional groups instead of the usual alkyl or aryl substituents. These functional groups promote an intramolecular condensation (cyclization) of intermediate ketone **II** or aldehyde **III** formed as a result of the above two transformations. Recently, our group reported a novel metal-free transannulation reaction of *N*,3-diaryloxirane-2-carboxamides (AOCAs) involving a one-pot acid-catalyzed Meinwald rearrangement and intramolecular Friedel–Crafts alkylation processes allowing to synthesize various 3-



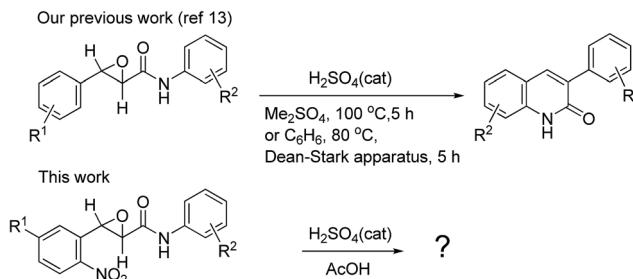
Scheme 1 Possible rearrangements of the trisubstituted oxiranes.

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† Electronic supplementary information (ESI) available: 1D and 2D NMR spectra and MALDI mass-spectra of the products **3a–h** and **4a–h**; crystallographic data for **4a,b** (CIF), description of quantum chemical computational setup and comparison of X-ray and DFT computed structural parameters of **4a**. CCDC 1417946 (for **4a**) and 1015263 (for **4b**). For ESI and crystallographic data in CIF or other electronic format see DOI: 10.1039/c6ra02586b





Scheme 2 Our previous work and this work.

arylquinolin-2(1*H*)-ones in high yields (Scheme 2).¹³ This novel approach features not only a metal-free bond formation but also an exclusive 1,2-aryl migration.

During our studies on the ring-opening/ring-closure reactions of AOCAs, we attempted to use *N*-aryl-3-(2-nitroaryl)oxirane-2-carboxamides (obtained from 2-nitrobenzaldehydes and 2-chloro-*N*-arylacetamides) with the aim of expanding the scope of the reaction. We found that compounds with a newly formed oxalamide chain were obtained instead of expected 3-(2-nitroaryl)quinolin-2(1*H*)-ones when the reactions were carried out in refluxing AcOH in the presence of H₂SO₄. As far as we know, there has been no report on the synthesis of unsymmetrical oxalamides *via* the rearrangement yet. Herein, we report this novel acid-catalyzed rearrangement of AOCAs in AcOH, which proceeds through a cascade of the ring-opening/ring-closure/ring-closure/ring-opening processes.

The salient features of our method are as follows: (1) a variety of aldehydes **1** and chloroacetamides **2** are readily available and the rapid synthesis of **3** with diverse substitution patterns are possible; (2) only two steps are necessary beginning with the starting materials to the products **4**; (3) the facile isolation of **3** and **4** are accomplished by a simple aqueous workup.

Results and discussion

The procedure of the Darzens condensation is the same as that described for AOCAs,¹³ except that only 2-nitrobenzaldehydes instead of variously substituted aromatic aldehydes with chloroacetamides were used and the reactions were carried out at room temperature for 7 h. The mixtures of *cis*- and *trans*-isomers of 3-(2-nitroaryl)oxirane-2-carboxamides (**3**) with the predominance of the *trans*-isomer were easily purified from the *cis*-isomer by washing with ether (Table 1).

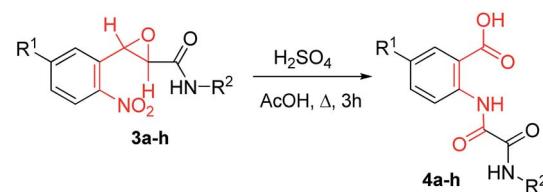
The structures of **3a–h** were proved by variety of 1D/2D NMR correlation methods (see ESI†).¹⁵ First of all to clarify the optimal reaction conditions we examined the rearrangement of *trans*-3-(2-nitrophenyl)-*N*-phenyloxirane-2-carboxamide (**3a**). After a brief survey of the reaction conditions, we have found that the product **4a** is obtained in almost quantitative yield at reflux for 3 h in AcOH with 1 equiv. of H₂SO₄ (Table 2, entry 1). The reflux of **3a** in both MeCN (with 1 equiv. H₂SO₄) and AcOH for 3 h resulted in the mixtures containing 30 and 10% (determined by ¹H NMR) of the desired product **4a**, respectively. Further optimization of the reaction conditions was carried out

Table 1 Synthesis of 3-(2-nitroaryl)oxirane-2-carboxamides

Entry	1	R ¹	2^a	R ²	Product	<i>trans/cis^b</i>	Yield _{trans,^c} %
1	1a	H	2a	Ph	3a^d	1/0.12	87
2	1a	H	2b	4-BrC ₆ H ₄	3b	1/0.14	85
3	1a	H	2c	3-MeC ₆ H ₄	3c	1/0.19	76
4	1a	H	2d	4-MeOC ₆ H ₄	3d	1/0.29	62
5	1a	H	2e	3-NO ₂ C ₆ H ₄	3e	1/0.07	91
6	1a	H	2f	4-CO ₂ EtC ₆ H ₄	3f	1/0.09	89
7	1b	Cl	2a	Ph	3g	1/0.30	61
8	1a	H	2g	H	3h	1/0.00	100

^a 2-Chloro-*N*-arylacetamides **2b–f** were obtained on a 0.1 mol scale at 0–15 °C by reacting chloroacetylchloride with an equimolar amounts of corresponding aniline and Et₃N. The compounds **2a** and **2g** are commercially available. ^b Ratio was determined by ¹H NMR of the crude products. ^c Yields refer to isolated *trans*-isomers of **3**. ^d *cis*-Isomer of this compound was obtained early.¹⁴

with *trans*-3-(2-nitrophenyl)oxirane-2-carboxamide (**3h**). The reflux of **3h** in H₂O with 1 equiv. H₂SO₄ for 5 h or its storage at room temp in AcOH with 1 equiv. of H₂SO₄ for 24 h gave 57 and 75% of the product **4h**, respectively. However, an almost quantitative yield of the rearrangement product was achieved when 1 equiv. of H₂SO₄ was used in boiling AcOH for 3 h (entry 8). The latter condition was used for the rearrangement of all the compounds **3**. The rearrangement proceeds equally well with the compounds **3** containing various substituents in an anilide moiety, no matter whether it is a strong electron donating (entry 4) or a strong electron withdrawing (entry 5) group. Interestingly, under the rearrangement conditions the

Table 2 Synthesis of *N*-(2-carboxaryl)oxalamides

Entry	3	R ¹	R ²	Product	Yield, ^a %
1	3a	H	Ph	4a	97
2	3b	H	4-BrC ₆ H ₄	4b	100
3	3c	H	3-MeC ₆ H ₄	4c	92
4	3d	H	4-MeOC ₆ H ₄	4d	95
5	3e	H	3-NO ₂ C ₆ H ₄	4e	96
6	3f	H	4-CO ₂ EtC ₆ H ₄	4f	97
7	3g	5-Cl	Ph	4g	98
8	3h	H	H	4h	97

^a Yields refer to isolated products.



ester group (entry 6) is not subjected to hydrolysis and it can be used in further transformations. The presence of the chlorine atom with the $-I$ and $+M$ electronic effects in the aldehyde component does not influence the yield of the rearrangement product (entry 7).

To our delight 3-(2-nitrophenyl)oxirane-2-carboxamides undergo the rearrangement with the formation of compounds which can be considered both as anthranilic acid and as unsymmetrical oxalamide derivatives. Anthranilic acid derivatives are potential anticancer agents¹⁶ and the ligands for farnesoid X receptor.¹⁷ Oxalamides also represent a key framework of many bioactive compounds.¹⁸ They have been developed as acetylcholine esterase inhibitors,¹⁹ C5a inhibitors,²⁰ nitric oxide synthase inhibitors,²¹ anti-HIV agents,²² antiepileptic drugs,²³ HIV integrase inhibitors,²⁴ HIV-1 proteas inhibitors,²⁵ cephalosporin bactericides²⁶ and chemotherapeutic agents.²⁷ Considering the well documented medicinal utility of anthranilic acid and oxalamide derivatives, these tethered combinations of the two scaffolds afford new opportunities to probe their biological activity.

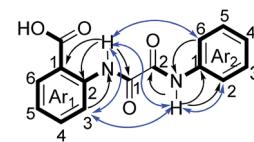
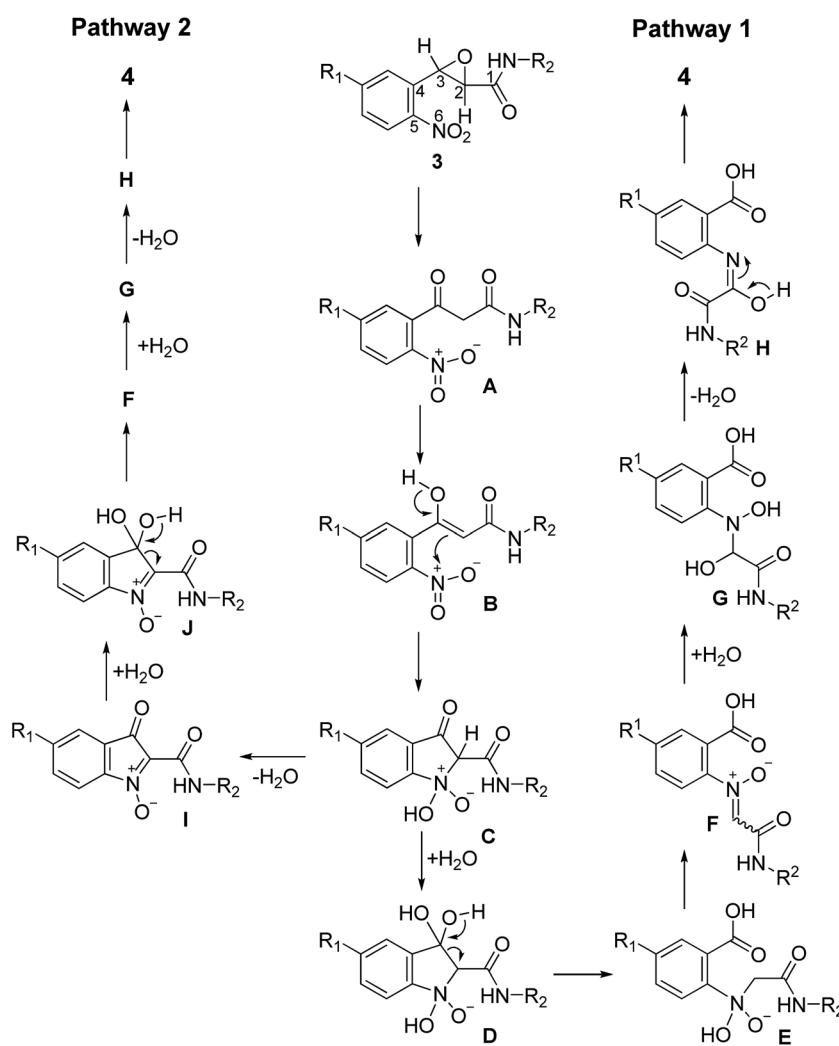


Fig. 1 Structure of **4a** with principle NMR correlations (^1H – ^{13}C / ^1H – ^{15}N HMBCs – black arrow, NOEs – blue arrow).

Based on above results and literature reports,^{3a,4,28a–c} a plausible mechanism for the rearrangement was proposed. First, the process was believed to proceed through the classical Meinwald rearrangement (Scheme 1, path *a*) of 3-(2-nitrophenyl)oxirane-2-carboxamides with the cleavage of the C2–O bond in its initial stage (**3** → **A**). The resulting ketone **A** bearing an active α -methylene group undergoes a new rearrangement according to the mechanism of the known Baeyer–Drewson indigo synthesis^{28a–c} with the formation of intermediate **C**, which is further subjected to acid-catalyzed ring opening (either **C** → **D** → **E** through pathway 1 or **C** → **I** → **J** → **F** through pathway 2)



Scheme 3 Proposed mechanisms of the rearrangement.



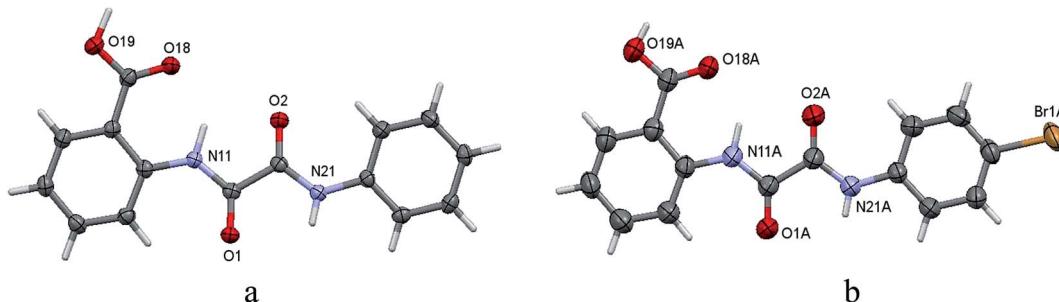
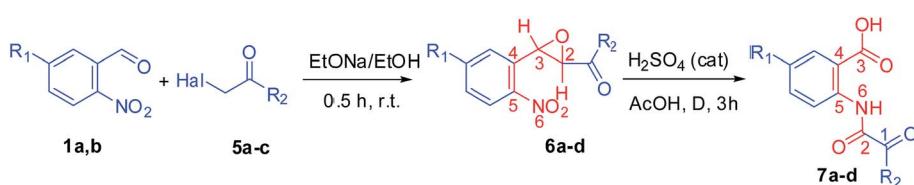


Fig. 2 ORTEP plot of compounds **4a** (a) and **4b** (b) partial numbering scheme. Displacement ellipsoids are drawn at the 50% probability level. H-atoms are represented in stick mode for clarity.

Table 3 Synthesis of 2-(2-oxo-2-phenylacetamido)benzoic acids **7a–d**



Entry	1	R^1	5^{a,b}	Hal	R^2	Product 6	Product 7	Yield, ^c %
1	1a	H	5a	Cl	4-MeC ₆ H ₄	6a	7a	75
2	1a	H	5b	Cl	4-MeOC ₆ H ₄	6b	7b	81
3	1a	H	5c	Br	3-NO ₂ C ₆ H ₄	6c	7c	86
4	1b	Cl	5a	Cl	4-MeC ₆ H ₄	6d	7d	89

^a (Chloromethyl)arylketones **5a,b** were obtained on a 0.1 mol scale at 10–20 °C by reaction of 1 equiv. chloroacetylchloride with toluene (100 mL) and anisole (100 mL), respectively, with the use of 1.5 equiv. AlCl_3 . ^b (Bromomethyl)arylketone **5c** was obtained on a 0.1 mol scale at 50–55 °C by reaction of 1 equiv. 1-(3-nitrophenyl)ethanone with 1 equiv. bromine in ethanol (100 mL). ^c Yields refer to isolated products.

involving hydration and dehydration processes (**E** → **F** → **G** → **H** → **4** through pathway 1 and **F** → **G** → **H** → **4** through pathway 2) with the participation of nitro-group and α -methylene functionalities. As a result, the reduction of the nitro-group and the transformation of the C3 atom of epoxide to the carboxylic functionality occurs (Scheme 3).

The structures of **4a–h** were established unambiguously by various 1D/2D NMR correlation methods.¹⁵ First, the proton spin systems of the Ar1 and Ar2 moieties were identified by COSY/TOCSY methods. After that, the structures of both halves up to carbonyl groups (C₁ and C₂) were established (boldfaced on Fig. 1) from the ¹H–¹³C and ¹H–¹⁵N HSQC/HMBC connectivities. Finally, both halves were linked into a single whole on the basis of the NOEs between protons of these two fragments:

Structures of the compounds **4a,b** were further confirmed by single-crystal X-ray analyses (Fig. 2).

It should be pointed out that a series of synthetic methods for oxalamides have been described in the past decades.²⁹ However, only five examples of the synthetic methods for unsymmetrical oxalamides are known. The first is traditional and based on the condensation of corresponding carboxylic acids with amines, which needs either activating agents or conversion into more reactive derivatives.³⁰ The next three

methods include the direct amidation of isocyanates,³¹ α -keto benzotriazole³² and trichloropyruvamides with amines.³³ The fifth method is a novel one and is based on green H_2O_2 -promoted oxidative amidation of 2-oxoaldehydes with amines.³⁴ Nevertheless, these methods have several drawbacks, such as harsh conditions, expensive reagents, poor atom-efficiency and limited substrate scope. Method proposed in this study demonstrated a new, efficient and metal-free synthesis of unsymmetrical oxalamides *via* novel rearrangement of easily available 3-(2-nitroaryl)oxirane-2-carboxamides.

With this result in hand, we proceeded with the study of the scope of the rearrangement. As can be seen from the suggested mechanism of the rearrangement (Scheme 3) the transformation of 3-(2-nitroaryl)oxirane-2-carboxamides to the oxalamides involves the 3-(2-nitroaryl)oxirane fragment only. Thus, seeking to expand accessible skeletal diversity using the same reaction conditions, we anticipated that use of (3-(2-nitroaryl)oxiran-2-yl)(aryl)methanones (**6a–d**) with the same necessary fragment would facilitate access to 2-(2-oxo-2-arylacetamido)benzoic acids (**7a–d**). Indeed the refluxing of oxiranes **6a–d** in AcOH in the presence of catalytic amount of H_2SO_4 provides the desired mono-oxalamides **7a–d** in good yields. Moreover, the reaction of 2-nitrobenzaldehyde (**1a**) and 5-chloro-2-



nitrobenzaldehyde (**1b**) with 2-chloro-1-(4-tolyl)ethanone (**5a**), 2-chloro-1-(4-methoxyphenyl)ethanone (**5b**) and 2-chloro-1-(3-nitrophenyl)ethanone (**5c**) under the Darzens condensation condition proceeded smoothly, and (3-(2-nitrophenyl)oxiran-2-yl)(aryl)methanones **6a–d** were obtained in quantitative yields (Table 3). It should be pointed out that in this case, in contrast to the reactions of chloroacetanilides **2**,¹³ the process proceeds with high stereoselectivity with the formation of only *trans*-isomers of oxiranes **6a–d** as the only products.

The structures of all compounds were proved by variety of 1D/2D NMR correlation methods (see ESI†).¹⁵

Conclusion

In conclusion, we have discovered a new rearrangement of 3-(2-nitrophenyl)-oxirane-2-carboxamides proceeding in boiling AcOH in the presence of H₂SO₄. The rearrangement quantitatively produces the *N*-(2-carboxyaryl)oxalamides as a result of cascade processes involving (a) the classical Meinwald rearrangement in its initial stage with the formation of ketone bearing an active α -methylene group, (b) transformation of carbonyl group of the ketone to the carboxylic functionality, (c) migration of active α -methylene group to the nitrogen atom of already reduced nitro group. The simple reaction conditions offer a potential for employing this method in the synthesis of complex molecules. It is anticipated that this methodology will have versatile applications in the practical syntheses of biologically important pharmaceutical molecules with anthranilic acid and oxalamide moieties. The methodology is applicable to synthesis of *N*-(2-carboxyphenyl)aryloxalmonoamides from (3-(2-nitrophenyl)oxiran-2-yl)(aryl)methanones. Further extension of the reaction scope and the synthetic applications of this methodology are in progress at our laboratory.

Experimental section

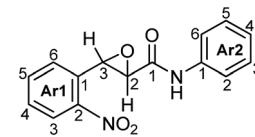
General methods

All reagents and solvents were used as purchased, without further purification melting points were determined on a hot-stage apparatus. Infrared (IR) spectra samples in Nujol were recorded on a FT-IR spectrometer Bruker Vector-22 in the 400–4000 cm^{−1} range at optical resolution of 4 cm^{−1}. The high resolution MALDI mass-spectra were obtained on UltraFlex III TOF/TOF instrument in positive reflectron mode; 2,5-DHB and *p*-NA were used as matrix and PEG-400 was used for calibration of accurate masses. All NMR experiments were performed with 600, 500 and 400 MHz (600 MHz for ¹H NMR; 150.9, 125 and 100.6 MHz for ¹³C NMR; 60.8 and 50.7 MHz for ¹⁵N NMR) spectrometers equipped with 5 mm diameter gradient inverse broad band probehead and a pulsed gradient unit capable of producing magnetic field pulse gradients in the z-direction of 53.5 G cm^{−1}. NMR experiments were carried out at 303 K. DPFGROE³⁵ and TOCSY spectra were obtained using a Hermite-shaped pulse for selective excitation. Chemical shifts (δ in ppm) are referred to the solvent DMSO-*d*₆ (δ = 2.49 ppm for ¹H and 39.5 ppm for ¹³C NMR), to external CD₃NO₂ (380.2 ppm) for ¹⁵N NMR spectra (conversion factor to NH₃: −380.2 ppm). ¹H–¹H

coupling constants were computed according to Bally & Rablen's recommendations.³⁶ First the geometry was optimized at the B3LYP/6-31G(d) level. Then NMR single-point calculation of the Fermi contact *J* values was run at the B3LYP/6-31G(d,p) level. These values were scaled then by a factor of 0.9117. The quantum chemical calculations were performed using a Gaussian 03 software package.³⁷

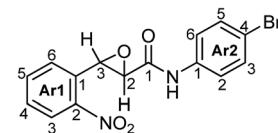
General procedure for the Darzens condensation. A solution of EtONa obtained when dissolving Na (0.73 g, 0.032 g-atom) in EtOH (20 mL) was added at room temperature to the stirred solution of 2-nitrobenzaldehyde (4.08 g, 0.027 mol) with the corresponding compound **2** (0.027 mol) in EtOH (70 mL). The stirring is continued for 7 h. Treatment of the reaction mixtures as is usual for previous cases¹³ led to the products **3**.

trans-3-(2-Nitrophenyl)-*N*-phenyloxirane-2-carboxamides (**3a**).



Tan powder (7.68 g, 0.027 mol, 87% yield): mp 180–181 °C; ¹H NMR (600 MHz, DMSO-*d*₆) δ 10.35 (s, 1H, NH), 8.20 (d, *J* = 8.2 Hz, 1H, H3-Ar1), 7.85 (dd, *J* = 7.6, 7.6 Hz, 1H, H5-Ar1), 7.65–7.69 (m, 3H, H2,6-Ar2; H4-Ar1), 7.59 (d, *J* = 7.9 Hz, 1H, H6-Ar1), 7.36 (dd, *J* = 7.9, 7.9 Hz, 2H, H3,5-Ar2), 7.12 (dd, *J* = 7.6, 7.2 Hz, 1H, H4-Ar2), 4.68 (d, *J* = 1.8 Hz, 1H, H3), 3.65 (d, *J* = 1.8 Hz, 1H, H2); ¹³C NMR (150.9 MHz, DMSO-*d*₆) δ 164.46 (C1), 147.46 (C2-Ar1), 138.23 (C1-Ar2), 134.66 (C5-Ar1), 132.19 (C1-Ar1), 129.50 (C4-Ar1), 128.78 (C3-Ar2), 126.73 (C6-Ar1), 124.65 (C3-Ar1), 123.91 (C4-Ar2), 119.45 (C2-Ar2), 57.37 (C2), 55.14 (C3); ¹⁵N NMR (60.8 MHz, DMSO-*d*₆) δ 371.6 (NO₂), 132.3 (NH); IR (nujol): ν 3279, 1674, 1604, 1553, 1525 cm^{−1}; HRMS (MALDI) calcd for C₁₅H₁₂N₂O₄ [M + Cs]⁺ 416.9846, found 416.9843; anal. calcd for C₁₅H₁₂N₂O₄: C, 63.38; H, 4.25; N, 9.85; found: C, 63.42; H, 4.13; N, 9.92.

trans-*N*-(4-Bromophenyl)-3-(2-nitrophenyl)oxirane-2-carboxamide (**3b**).

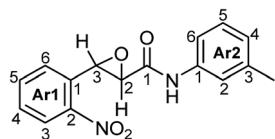


Tan powder (8.33 g, 0.023 mol, 85% yield): mp 191–192 °C; ¹H NMR (600 MHz, DMSO-*d*₆) δ 10.50 (s, 1H, NH), 8.19 (dd, *J* = 8.2, 1.0 Hz, 1H, H3-Ar1), 7.85 (dd, *J* = 7.7, 7.2 Hz, 1H, H5-Ar1), 7.67 (ddd, *J* = 8.2, 7.7, 1.0 Hz, 1H, H4-Ar1), 7.64 (d, *J* = 8.7 Hz, 2H, H2,4-Ar2), 7.58 (d, *J* = 7.7 Hz, 1H, H6-Ar1), 7.54 (d, *J* = 8.7 Hz, 2H, H3,5-Ar2), 4.68 (d, *J* = 1.8 Hz, 1H, H3), 3.64 (d, *J* = 1.8 Hz, 1H, H2); ¹³C NMR (100.6 MHz, DMSO-*d*₆) δ 164.74 (C1), 147.45 (C2-Ar1), 137.59 (C1-Ar2), 134.69 (C5-Ar1), 132.09 (C1-Ar1), 131.64 (C4-Ar2), 129.56 (C4-Ar1), 126.73 (C6-Ar1), 124.68 (C3-Ar1), 121.43 (C2-Ar2), 115.61 (C4-Ar2), 57.36 (C2), 55.27 (C3); ¹⁵N NMR (60.8 MHz, DMSO-*d*₆) δ 371.5 (NO₂), 131.2 (NH). IR (nujol): ν 3362, 1691, 1591, 1521 cm^{−1}; HRMS (MALDI) calcd for C₁₅H₁₁BrN₂O₄ [M + Cs]⁺ 494.8951; 496.8932, found 494.8946;



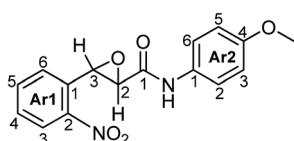
496.8938; anal. calcd for $C_{15}H_{11}BrN_2O_4$: C, 49.61; H, 3.05; Br, 22.00; N, 7.71; found: C, 49.84; H, 2.99; Br, 21.84; N, 7.87.

trans-3-(2-Nitrophenyl)-N-3-tolylloxirane-2-carboxamide (3c).



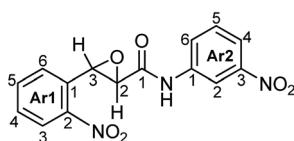
Tan powder (6.12 g, 0.021 mol, 76% yield): mp 140–143 °C; 1H NMR (600 MHz, DMSO- d_6): δ 10.29 (s, 1H, NH); 8.21 (dd, J = 8.2, 1.1 Hz, 1H, H3-Ar1), 7.86 (ddd, J = 7.5, 7.2, 1.1 Hz, 1H, H5-Ar1), 7.68 (ddd, J = 7.9, 7.8, 1.4 Hz, 1H, H4-Ar1), 7.59 (d, J = 7.6 Hz, 1H, H2-Ar1), 7.51 (br.s, 1H, H2-Ar2), 7.45 (br.d, J = 8.2 Hz, 1H, H6-Ar2), 7.24 (dd, J = 7.8, 7.7 Hz, 1H, H5-Ar2), 6.94 (d, J = 7.4 Hz, 1H, H4-Ar2), 4.68 (d, J = 2.0 Hz, 1H, H3), 3.65 (d, J = 2.0 Hz, 1H, H2), 2.31 (s, 3H, CH₃); ^{13}C NMR (125.8 MHz, DMSO- d_6) δ 164.39 (C1), 147.45 (C2-Ar1), 138.17 (C3-Ar2), 138.05 (C1-Ar2), 134.69 (C5-Ar1), 132.24 (C1-Ar1), 129.51 (C4-Ar1), 128.64 (C5-Ar2), 126.73 (C6-Ar1), 124.67 (C4-Ar2), 124.62 (C3-Ar1), 119.96 (C2-Ar2), 116.65 (C6-Ar2), 57.38 (C2), 55.14 (C3), 21.09 (Me); ^{15}N NMR (60.8 MHz, DMSO- d_6) δ 371.6 (NO₂), 132.5 (NH); IR (nujol): ν 3241, 1673, 1535, 1522 cm⁻¹; HRMS (MALDI) calcd for $C_{16}H_{14}N_2O_4$ [M + Cs]⁺ 431.0003, found 430.9994; anal. calcd for $C_{16}H_{14}N_2O_4$: C, 64.42; H, 4.73; N, 9.39; found: C, 64.86; H, 4.72; N, 9.41.

trans-N-(4-Methoxyphenyl)-3-(2-nitrophenyl)oxirane-2-carboxamide (3d).



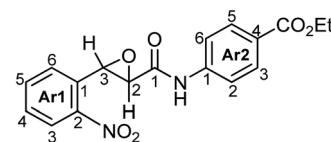
Brown powder (5.34 g, 0.017 mol, 62% yield): mp 168–169 °C; 1H NMR (600 MHz, DMSO- d_6): δ 10.23 (s, 1H, NH), 8.21 (dd, J = 8.2, 1.1 Hz, 1H, H3-Ar1), 7.86 (ddd, J = 7.6, 7.6, 1.0 Hz, 1H, H5-Ar1), 7.68 (ddd, J = 7.8, 7.8, 1.3 Hz, 1H, H4-Ar1), 7.49–7.60 (m, 3H, H2,4-Ar2; H6-Ar1), 6.93 (d, J = 9.1 Hz, 2H, H3,5-Ar2), 4.68 (d, J = 1.9 Hz, 1H, H3), 3.75 (s, 3H, OCH₃), 3.61 (d, J = 1.9 Hz, 1H, H2); ^{13}C NMR (125.8 MHz, DMSO- d_6) δ 163.98 (C1), 155.67 (C4-Ar2), 147.47 (C2-Ar1), 134.69 (C5-Ar1), 132.28 (C1-Ar1), 131.40 (C1-Ar2), 129.51 (C4-Ar1), 126.75 (C6-Ar1), 124.68 (C3-Ar1), 121.10 (C2-Ar2), 113.94 (C3-Ar2), 57.44 (C2), 55.17 (OMe), 55.10 (C3); ^{15}N NMR (60.8 MHz, DMSO- d_6) δ 371.8 (NO₂-Ar1), 130.6 (NH); IR (nujol): ν 3268, 1664, 1608, 1595, 1556, 1514 cm⁻¹; HRMS (MALDI) calcd for $C_{16}H_{14}N_2O_5$ [M + Na]⁺ 337.0795, found 337.0814; anal. calcd for $C_{16}H_{14}N_2O_5$: C, 61.14; H, 4.50; N, 8.91. Found: C, 61.53; H, 4.48; N, 8.93.

trans-N-(3-Nitrophenyl)-3-(2-nitrophenyl)oxirane-2-carboxamide (3e).



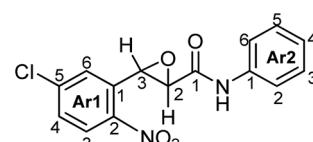
Tan powder (8.09 g, 0.025 mol, 91% yield): mp 177 °C; 1H NMR (600 MHz, DMSO- d_6): δ 10.83 (s, 1H, NH), 8.70 (dd, J = 2.2, 2.2 Hz, 1H, H2-Ar2), 8.21 (dd, J = 8.3, 1.0 Hz, 1H, H3-Ar1), 8.01 (dd, J = 8.2, 1.5 Hz, 1H, H6-Ar2), 7.98 (dd, J = 8.2, 1.3 Hz, 1H, H4-Ar2), 7.86 (dd, J = 7.7, 7.3 Hz, 1H, H5-Ar1), 7.68 (dd, J = 7.7, 1.4 Hz, 1H, H4-Ar1), 7.66 (dd, J = 8.2, 8.2 Hz, 1H, H5-Ar2), 7.60 (d, J = 7.7 Hz, 1H, H6-Ar1), 4.73 (d, J = 2.0 Hz, 1H, H3), 3.70 (d, J = 2.0 Hz, 1H, H2); ^{13}C NMR (150.9 MHz, DMSO- d_6) δ 165.44 (C1), 147.95 (C3-Ar2), 147.44 (C2-Ar1), 139.27 (C1-Ar2), 134.70 (C5-Ar1), 131.90 (C1-Ar1), 130.29 (C5-Ar2), 129.62 (C4-Ar1), 126.76 (C6-Ar1), 125.50 (C6-Ar2), 124.68 (C3-Ar1), 118.47 (C4-Ar2), 113.72 (C2-Ar2), 57.26 (C2), 55.48 (C3); ^{15}N NMR (60.8 MHz, DMSO- d_6) δ 371.4 (NO₂-Ar1), 370.3 (NO₂-Ar2), 130.7 (NH); IR (nujol): ν 3241, 1673, 1535, 1522 cm⁻¹; HRMS (MALDI) calcd for $C_{15}H_{11}N_3O_6$ [M + Na]⁺ 352.0540, found 352.0559; anal. calcd for $C_{15}H_{11}N_3O_6$: C, 54.72; H, 3.37; N, 12.76. Found: C, 54.69; H, 3.28; N, 12.78.

trans-3-(2-Nitrophenyl)-N-(4-ethylcarboxyphenyl)oxirane-2-carboxamide (3f).



White powder (8.56 g, 0.024 mol, 89% yield): mp 180–182 °C; 1H NMR (600 MHz, DMSO- d_6): δ 10.86 (br.s, 1H, NH), 8.21 (d, J = 8.2 Hz, 1H, H3-Ar1), 7.96 (d, J = 8.5 Hz, 2H, H3,5-Ar2), 7.86 (dd, J = 7.6, 7.6 Hz, 1H, H5-Ar1), 7.83 (d, J = 8.5 Hz, 2H, H2,6-Ar2), 7.68 (dd, J = 8.0, 7.7 Hz, 1H, H4-Ar1), 7.59 (d, J = 7.2 Hz, 1H, H6-Ar1), 4.71 (d, J = 1.7 Hz, 1H, H3), 3.75 (d, J = 1.7 Hz, 1H, H2), 4.30 (q, J = 7.1 Hz, 2H, OCH₂CH₃), 1.33 (t, J = 7.1 Hz, 3H, OCH₂CH₃); ^{13}C NMR (125.8 MHz, DMSO- d_6) δ 165.21 (CO₂Et), 165.17 (C1), 147.46 (C2-Ar1), 142.57 (C1-Ar2), 134.72 (C5-Ar1), 132.11 (C1-Ar1), 130.24 (C3-Ar2), 129.59 (C4-Ar1), 126.78 (C6-Ar1), 124.93 (C4-Ar2), 124.70 (C3-Ar1), 118.94 (C2-Ar2), 60.46 (OCH₂CH₃), 57.28 (C2), 55.37 (C3), 14.15 (OCH₂CH₃); ^{15}N NMR (60.8 MHz, DMSO- d_6) δ 371.7 (NO₂), 134.1 (NH); IR (nujol): ν 3360, 1711, 1697, 1608, 1595, 1522 cm⁻¹; HRMS (MALDI) calcd for $C_{18}H_{16}N_2O_6$ [M + Cs]⁺ 489.0057, found 489.0055; anal. calcd for $C_{18}H_{16}N_2O_6$: C, 60.67; H, 4.53; N, 7.86. Found: C, 60.68; H, 4.48; N, 7.93.

trans-3-(2-Nitro-5-chlorophenyl)-N-phenyloxirane-2-carboxamide (3g).

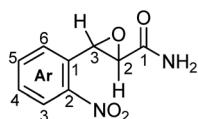


White powder (5.25 g, 0.016 mol, 61% yield): mp 156–158 °C; 1H NMR (600 MHz, DMSO- d_6): δ 10.38 (s, 1H, NH), 8.25 (d, J = 8.8 Hz, 1H, H3-Ar1), 7.75 (dd, J = 8.8, 2.4 Hz, 1H, H4-Ar1), 7.67 (d, J = 8.4 Hz, 2H, H2,6-Ar2), 7.56 (d, J = 2.4 Hz, 1H, H6-Ar1), 7.38 (dd, J = 8.4, 7.5 Hz, 2H, H3,5-Ar2), 7.13 (ddd, J = 7.5, 7.4, 1.0 Hz, 1H, H4-Ar2), 4.72 (d, J = 2.0 Hz, 1H, H3), 3.72 (d, J = 2.0 Hz, 1H,



H2); ^{13}C NMR (125.8 MHz, DMSO- d_6) δ 164.23 (C1), 146.12 (C2-Ar1), 139.53 (C5-Ar1), 138.19 (C1-Ar2), 134.65 (C1-Ar1), 129.46 (C4-Ar1), 128.82 (C3-Ar2), 126.90 (C3-Ar1), 126.53 (C6-Ar1), 123.99 (C4-Ar2), 119.50 (C2-Ar2), 57.34 (C2), 54.84 (C3); ^{15}N NMR (60.8 MHz, DMSO- d_6) δ 368.9 (NO₂), 132.6 (NH); IR (nujol): ν 3368, 1693, 1600, 1571, 1533 cm⁻¹; HRMS (MALDI) calcd for C₁₅H₁₁ClN₂O₄ [M + Na]⁺ 341.0299, found 341.0308; anal. calcd for C₁₅H₁₁ClN₂O₄: C, 56.53; H, 3.48; Cl, 11.12; N, 8.79. Found: C, 56.62; H, 3.32; Cl, 11.18; N, 8.92.

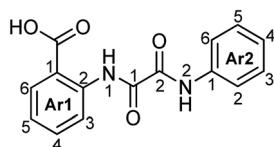
trans-3-(2-Nitrophenyl)oxirane-2-carboxamide (3h).



Light yellow powder (5.62 g, 0.027 mol, 100% yield); mp 209–210 °C; ^1H NMR (600 MHz, DMSO- d_6) δ 8.18 (dd, J = 8.1, 1.0 Hz, 1H, H3-Ar1), 7.82 (ddd, J = 7.8, 7.8, 1.0 Hz, 1H, H5-Ar1), 7.69 (br.s, 1H, NH₂); 7.65 (ddd, J = 8.1, 7.8, 1.2 Hz, 1H, H4-Ar1), 7.48 (br.s, 1H, NH₂); 7.53 (d, J = 7.7 Hz, 1H, H6-Ar1), 4.52 (d, J = 1.9 Hz, 1H, H3), 3.39 (d, J = 1.9 Hz, 1H, H2); ^{13}C NMR (125.8 MHz, DMSO- d_6) δ 168.07 (C1), 147.50 (C2-Ar), 134.68 (C5-Ar), 132.43 (C1-Ar), 129.46 (C4-Ar), 126.73 (C6-Ar), 124.68 (C3-Ar), 57.05 (C2), 54.91 (C3); ^{15}N NMR (50.7 MHz, DMSO- d_6) δ 371.8 (NO₂), 106.4 (NH₂); IR (nujol): ν 3368, 3185, 1665, 1526 cm⁻¹; HRMS (MALDI) calcd for C₉H₈N₂O₄ [M + Cs]⁺ 340.9533, found 340.9536; anal. calcd for C₉H₈N₂O₄: C, 51.93; H, 3.87; N, 13.46. Found: C, 52.02; H, 3.85; N, 13.43.

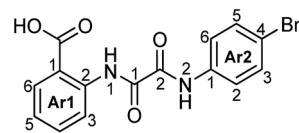
General procedure for the rearrangement. Concd H₂SO₄ (0.1 mL) was added to the solution of 3 (1.67 mmol) in AcOH (5 mL). The reaction mixture was boiled for 3 h and then poured into water. The precipitate thus formed was collected by filtration, washed with water and dried.

*N*¹-(2-Carboxyphenyl)-*N*²-phenyloxalamide (4a).



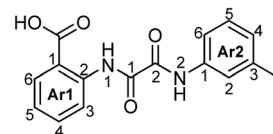
Light brown powder (0.46 g, 1.62 mmol, 97% yield); mp 227–228 °C; ^1H NMR (600 MHz, DMSO- d_6) δ 12.75 (s, 1H, N1H), 10.89 (s, 1H, N2H), 8.70 (ddd, J = 8.3, 8.3, 1.0 Hz, 1H, H3-Ar1), 8.08 (dd, J = 8.0, 1.5 Hz, 1H, H6-Ar1), 7.88 (dd, J = 8.7, 1.1 Hz, 2H, H2,6-Ar2), 7.71 (ddd, J = 7.9, 7.9, 1.7 Hz, 1H, H4-Ar1), 7.39 (ddd, J = 8.7, 7.6, 1.1 Hz, 2H, H3,5-Ar2), 7.29 (ddd, J = 7.6, 7.6, 1.1 Hz, 1H, H5-Ar1), 7.18 (ddd, J = 7.4, 7.4, 1.0 Hz, 1H, H4-Ar2); ^{13}C NMR (125.8 MHz, DMSO- d_6) δ 168.84 (CO₂H), 158.29 (2C=O), 158.03 (1C=O), 139.16 (C2-Ar1), 137.41 (C1-Ar2), 134.19 (C4-Ar1), 131.43 (C6-Ar1), 128.65 (C3-Ar2), 124.72 (C4-Ar2), 123.92 (C5-Ar1), 120.60 (C2-Ar2), 119.53 (C3-Ar1), 117.39 (C1-Ar1); ^{15}N NMR (60.8 MHz, DMSO- d_6) δ 126.8 (N2), 121.7 (N1); IR (nujol): ν 3329, 3179, 1678, 1586, 1520 cm⁻¹; HRMS (MALDI) calcd for C₁₅H₁₂N₂O₄ [M + Na]⁺ 307.0689, found 307.0690; anal. calcd for C₁₅H₁₂N₂O₄: C, 63.38; H, 4.25; N, 9.85. Found: C, 63.34; H, 4.31; N, 9.93.

*N*¹-(2-Carboxyphenyl)-*N*²-(4-bromophenyl)oxalamide (4b).



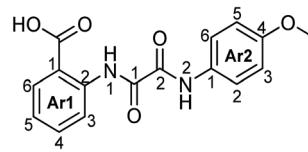
Grey powder (0.61 g, 1.67 mmol, 100% yield); mp 281–283 °C; ^1H NMR (600 MHz, DMSO- d_6) δ 12.72 (s, 1H, N1H), 11.04 (s, 1H, N2H), 8.68 (d, J = 8.2 Hz, 1H, H3-Ar1), 8.07 (dd, J = 8.2, 1.5 Hz, 1H, H6-Ar1), 7.85 (d, J = 8.7 Hz, 2H, H2,6-Ar2), 7.71 (ddd, J = 7.9, 7.7, 1.5 Hz, 1H, H4-Ar1), 7.57 (d, J = 8.7 Hz, 2H, H3,5-Ar2), 7.28 (ddd, J = 7.7, 7.7, 1.0 Hz, 1H, H5-Ar1); ^{13}C NMR (150.9 MHz, DMSO- d_6) δ 168.82 (CO₂H), 158.18 (2C=O), 158.03 (1C=O), 139.10 (C2-Ar1), 136.84 (C1-Ar2), 134.20 (C4-Ar1), 131.49 (C3-Ar2), 131.41 (C6-Ar1), 123.95 (C5-Ar1), 122.54 (C2-Ar2), 119.54 (C3-Ar1), 117.32 (C1-Ar1), 116.65 (C4-Ar2); ^{15}N NMR (60.8 MHz, DMSO- d_6) δ 126.6 (N2), 122.2 (N1); IR (nujol): ν 3301, 3189, 1687, 1584, 1519 cm⁻¹; HRMS (MALDI) calcd for C₁₅H₁₁BrN₂O₄ [M + Na]⁺ 384.9794; 386.9776, found 384.9795; 386.9786; anal. calcd for C₁₅H₁₁BrN₂O₄: C, 49.61; H, 3.05; Br, 22.00; N, 7.71. Found: C, 49.38; H, 3.02; Br, 22.07; N, 7.69.

*N*¹-(2-Carboxyphenyl)-*N*²-(3-methylphenyl)oxalamide (4c).



Brown powder (0.46 g, 1.54 mmol, 92% yield); mp 203–205 °C; ^1H NMR (600 MHz, DMSO- d_6) δ 12.72 (s, 1H, N1H), 10.79 (s, 1H, N2H), 8.70 (d, J = 8.4, 1.0 Hz, 1H, H3-Ar1), 8.08 (dd, J = 7.9, 1.6 Hz, 1H, H6-Ar1), 7.73 (br.s, 1H, H2-Ar2), 7.72 (ddd, J = 7.8, 7.8, 1.7 Hz, 1H, H4-Ar1), 7.64 (br.d, J = 8.3 Hz, 1H, H6-Ar1), 7.29 (ddd, J = 7.9, 7.9, 1.2 Hz, 1H, H5-Ar1), 7.27 (dd, J = 7.8, 7.8 Hz, 1H, H5-Ar2), 7.00 (d, J = 7.6 Hz, 1H, H4-Ar2), 2.33 (s, 3H, CH₃); ^{13}C NMR (125.8 MHz, DMSO- d_6) δ 168.83 (CO₂H), 158.32 (2C=O), 157.95 (1C=O), 139.16 (C2-Ar1), 137.92 (C3-Ar2), 137.31 (C1-Ar2), 134.24 (C4-Ar1), 131.44 (C6-Ar1), 128.50 (C5-Ar2), 125.44 (C4-Ar2), 123.94 (C5-Ar1), 121.06 (C2-Ar2), 119.53 (C3-Ar1), 117.82 (C6-Ar2), 117.31 (C1-Ar1), 21.12 (Me); ^{15}N NMR (50.7 MHz, DMSO- d_6) δ 126.9 (N2), 121.8 (N1); IR ν (nujol): ν 3317, 3187, 1678, 1587, 1528 cm⁻¹; HRMS (MALDI) calcd for C₁₆H₁₄N₂O₄ [M + Cs]⁺ 431.0003, found 431.0014; anal. calcd for C₁₆H₁₄N₂O₄: C, 64.42; H, 4.73; N, 9.39. Found: C, 64.31; H, 4.68; N, 9.42.

*N*¹-(2-Carboxyphenyl)-*N*²-(4-methoxyphenyl)oxalamide (4d).

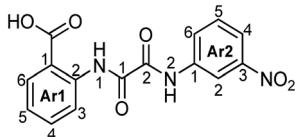


Grey powder (0.50 g, 1.59 mmol, 95% yield); mp 272–273 °C; ^1H NMR (600 MHz, DMSO- d_6) δ 12.70 (s, 1H, N1H), 10.78 (s, 1H, N2H), 8.69 (ddd, J = 8.4, 7.7, 1.0 Hz, 1H, H3-Ar1), 8.07 (dd, J = 8.2, 1.6 Hz, 1H, H6-Ar1), 7.78 (d, J = 9.1 Hz, 2H, H2,6-Ar2), 7.71



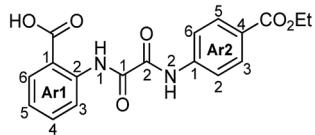
(ddd, $J = 7.9, 7.8, 1.6$ Hz, 1H, H4-Ar1), 7.28 (ddd, $J = 7.8, 7.4, 1.1$ Hz, 1H, H5-Ar1), 6.95 (d, $J = 9.1$ Hz, 2H, H3,5-Ar2), 3.76 (s, 3H, OCH₃); ¹³C NMR (125.8 MHz, DMSO-*d*₆) δ 168.91 (CO₂H), 158.56 (2C=O), 157.66 (1C=O), 156.34 (C4-Ar2), 139.24 (C2-Ar1), 134.31 (C4-Ar1), 131.51 (C6-Ar1), 130.54 (C1-Ar2), 124.01 (C5-Ar1), 122.15 (C2-Ar2), 119.63 (C3-Ar1), 117.37 (C1-Ar1), 113.91 (C3-Ar2), 55.27 (OMe); ¹⁵N NMR (50.7 MHz, DMSO-*d*₆) δ 125.6 (N2), 121.8 (N1); IR (nujol): ν 3336, 3218, 1683, 1585, 1525 cm⁻¹; HRMS (MALDI) calcd for C₁₆H₁₄N₂O₅ [M + Cs]⁺ 446.9952, found 446.9953; anal. calcd for C₁₆H₁₄N₂O₅: C, 61.14; H, 4.50; N, 8.91. Found: C, 60.99; H, 4.58; N, 8.97.

*N*¹-(2-Carboxyphenyl)-*N*²-(3-nitrophenyl)oxalamide (4e).



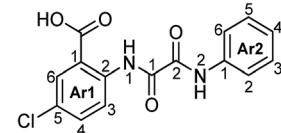
Brown powder (0.53, 1.61 mmol, 96% yield): mp 254–256 °C; ¹H NMR (600 MHz, DMSO-*d*₆) δ 12.90 (s, 1H, N1H), 11.40 (s, 1H, N2H), 8.90 (dd, $J = 2.0, 2.0$ Hz, 1H, H2-Ar2), 8.69 (d, $J = 8.2$ Hz, 1H, H3-Ar1), 8.27 (dd, $J = 7.9, 1.2$ Hz, 1H, H6-Ar2), 8.08 (dd, $J = 7.9, 1.2$ Hz, 1H, H6-Ar1), 8.02 (dd, $J = 7.9, 1.8$ Hz, 1H, H4-Ar2), 7.66–7.72 (m, 2H, H5-Ar2; H4-Ar1), 7.28 (dd, $J = 7.8, 7.2$ Hz, 1H, H5-Ar1); ¹³C NMR (150.9 MHz, DMSO-*d*₆) δ 168.93 (CO₂H), 158.76 (2C=O), 157.68 (1C=O), 147.87 (C3-Ar2), 139.05 (C2-Ar1), 138.71 (C1-Ar2), 134.02 (C4-Ar1), 131.43 (C6-Ar1), 130.09 (C5-Ar2), 126.68 (C6-Ar2), 123.97 (C5-Ar1), 119.51 (C3-Ar2), 119.17 (C4-Ar2), 117.80 (C1-Ar1), 114.81 (C2-Ar2); ¹⁵N NMR (60.8 MHz, DMSO-*d*₆) δ 370.1 (NO₂), 126.6 (N2), 122.2 (N1); IR (nujol): ν 3326, 3188, 1690, 1589, 1532 cm⁻¹; HRMS (MALDI) calcd for C₁₅H₁₁N₃O₆ [M + Na]⁺ 352.0540, found 352.0563; anal. calcd for C₁₅H₁₁N₃O₆: C, 54.72; H, 3.37; N, 12.76. Found: C, 54.82; H, 3.43; N, 12.68.

*N*¹-(2-Carboxyphenyl)-*N*²-(4-ethylcarboxyphenyl)oxalamide (4f).



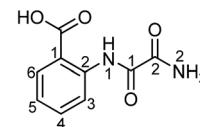
Light green powder (0.58 g, 1.63 mmol, 97% yield): mp 268–269 °C; ¹H NMR (600 MHz, DMSO-*d*₆) δ 12.78 (s, 1H, N1H), 11.18 (s, 1H, N2H), 8.68 (d, $J = 8.8$ Hz, 1H, H3-Ar1), 8.08 (d, $J = 7.7$ Hz, 1H, H6-Ar1), 8.04 (d, $J = 8.8$ Hz, 2H, H2,6-Ar2), 7.96 (d, $J = 8.8$ Hz, 2H, H3,5-Ar2), 7.70 (dd, $J = 7.7, 7.7$ Hz, 1H, H4-Ar1), 7.28 (dd, $J = 7.7, 7.7$ Hz, 1H, H5-Ar1), 4.30 (q, $J = 7.1$ Hz, 2H, OCH₂CH₃), 1.32 (t, $J = 7.1$ Hz, 3H, OCH₂CH₃); ¹³C NMR (150.9 MHz, DMSO-*d*₆) δ 168.90 (CO₂H), 165.18 (CO₂Et), 158.52 (2C=O), 157.88 (1C=O), 141.74 (C1-Ar2), 139.12 (C2-Ar1), 134.17 (C4-Ar1), 131.45 (C6-Ar1), 129.94 (C3-Ar2), 125.70 (C4-Ar2), 123.97 (C5-Ar1), 120.12 (C2-Ar2), 119.53 (C3-Ar1), 117.47 (C1-Ar1), 60.52 (OCH₂CH₃), 14.13 (OCH₂CH₃); ¹⁵N NMR (60.8 MHz, DMSO-*d*₆) δ 127.6 (N2), 122.0 (N1); IR (nujol): ν 3355, 3260, 1708, 1688, 1602, 1586, 1524, 1271, 761 cm⁻¹; HRMS (MALDI) calcd for C₁₈H₁₆N₂O₆ [M + 2Cs – H]⁺ 620.9034, found 620.9042; anal. calcd for C₁₈H₁₆N₂O₆: C, 60.67; H, 4.53; N, 7.86. Found: C, 60.59; H, 4.48; N, 7.97.

*N*¹-(4-Chloro-2-carboxyphenyl)-*N*²-phenyloxalamide (4g).



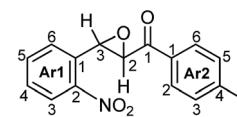
Brown powder (0.52 g, 1.63 mmol, 98% yield): mp 242–243 °C; ¹H NMR (600 MHz, DMSO-*d*₆) δ 12.84 (s, 1H, N1H), 10.89 (s, 1H, N2H), 8.70 (d, $J = 8.9$ Hz, 1H, H3-Ar1), 8.02 (d, $J = 2.6$ Hz, 1H, H6-Ar1), 7.86 (d, $J = 7.8$ Hz, 2H, H2,6-Ar2), 7.76 (dd, $J = 8.9, 2.6$ Hz, 1H, H4-Ar1), 7.39 (dd, $J = 8.1, 7.5$ Hz, 2H, H3,5-Ar2), 7.18 (dd, $J = 7.5, 7.4$ Hz, 1H, H4-Ar2); ¹³C NMR (100.9 MHz, DMSO-*d*₆) δ 167.59 (CO₂H), 158.34 (2C=O), 157.86 (1C=O), 137.98 (C2-Ar1), 137.37 (C1-Ar2), 133.58 (C4-Ar1), 130.61 (C6-Ar1), 128.64 (C3-Ar2), 127.51 (C5-Ar1), 124.74 (C4-Ar2), 121.29 (C3-Ar1), 120.61 (C2-Ar2), 119.71 (C1-Ar1); ¹⁵N NMR (50.7 MHz, DMSO-*d*₆) δ 127.0 (N2), 121.4 (N1); IR (nujol): ν 3331, 3166, 1679, 1579, 1505 cm⁻¹; HRMS (MALDI) calcd for C₁₅H₁₁ClN₂O₄ [M + Na]⁺ 341.0300, found 341.0328; anal. calcd for C₁₅H₁₁ClN₂O₄: C, 56.53; H, 3.48; Cl, 11.12; N, 8.79. Found: C, 56.59; H, 3.52; Cl, 11.07; N, 8.99.

N-(2-Carboxyphenyl)oxalamide (4h).



Violet powder (0.34 g, 1.62 mmol, 97% yield): mp 267–268 °C; ¹H NMR (600 MHz, DMSO-*d*₆) δ 12.51 (s, 1H, N1H), 8.65 (d, $J = 8.2$ Hz, 1H, H3-Ar), 8.36 (br.s, 1H, NH₂), 8.05 (br.s, 1H, NH₂), 8.04 (dd, $J = 8.0, 1.6$ Hz, 1H, H6-Ar), 7.68 (ddd, $J = 7.9, 7.9, 1.5$ Hz, 1H, H4-Ar), 7.25 (ddd, $J = 7.6, 7.6, 1.0$ Hz, 1H, H5-Ar); ¹³C NMR (125.8 MHz, DMSO-*d*₆) δ 168.66 (CO₂H), 161.57 (1C=O), 158.73 (2C=O), 139.19 (C2-Ar), 134.13 (C4-Ar), 131.38 (C6-Ar), 123.70 (C5-Ar), 119.47 (C3-Ar), 117.22 (C1-Ar); ¹⁵N NMR (50.7 MHz, DMSO-*d*₆) δ 103.5 (N2), 121.3 (N1); IR (nujol): ν 3538, 3465, 3321, 3168, 1729, 1683, 1592, 1537, 1272, 752 cm⁻¹; HRMS (MALDI) calcd for C₉H₈N₂O₄ [M + 2Cs – H]⁺ 472.8509, found 472.8491; anal. calcd for C₉H₈N₂O₄: C, 51.93; H, 3.87; N, 13.46. Found: C, 51.90; H, 3.89; N, 13.42.

Synthesis of (3-(2-nitroaryl)oxiran-2-yl)(aryl)methanones (6a–d). Synthesis of (3-(2-nitroaryl)oxiran-2-yl)(aryl)methanones (6a–d) was performed according to the general procedure for the Darzens condensation with the use of corresponding (halomethyl)arylketones (5a–c) instead of chloroacetamides (2a–g). In these cases as distinct from the synthesis of 3-(2-nitroaryl)oxiran-2-carboxamides (3a–h) the reactions are completed for 0.5 h. The products precipitated during the reaction and did not require any purification except washing with water (3 × 25 mL). *trans*-(3-(2-Nitrophenyl)oxiran-2-yl)(4-tolyl)methanone (6a).

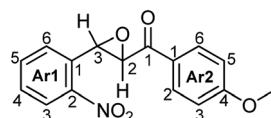


White powder (7.65 g, 0.027 mol, 100% yield): mp 154 °C; ¹H NMR (400 MHz, DMSO-*d*₆) δ 8.21 (dd, $J = 8.1, 1.0$ Hz, 1H, H3-



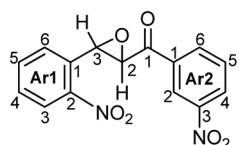
Ar1), 8.0 (d, J = 8.2 Hz, 2H, H3,5-Ar2), 7.88 (ddd, J = 7.6, 7.5, 0.8 Hz, 1H, H5-Ar1), 7.65–7.71 (m, 2H, H4,6-Ar1), 7.37 (d, J = 8.2 Hz, 2H, H2,6-Ar1), 4.74 (d, J = 2.1 Hz, 1H, H2), 4.56 (d, J = 2.1 Hz, 1H, H3), 2.41 (s, 3H, Me); ^{13}C NMR (100.6 MHz, DMSO- d_6) δ 192.14 (1C=O), 147.47 (C2-Ar1), 144.72 (C4-Ar2), 134.60 (C5-Ar1), 132.70 (C1-Ar2), 132.42 (C1-Ar1), 129.47 (C4-Ar1), 129.38 (C2-Ar2), 128.50 (C3-Ar2), 127.02 (C6-Ar1), 124.62 (C3-Ar1), 58.34 (C2), 57.15 (C3), 21.19 (Me). IR (nujol): ν 1682, 1522 cm^{-1} ; HRMS (MALDI) calcd for $\text{C}_{16}\text{H}_{13}\text{NO}_4$ [M + Cs]⁺ 415.9894, found 415.9895; anal. calcd for $\text{C}_{16}\text{H}_{13}\text{NO}_4$: C, 67.84; H, 4.63; N, 4.94; found: C, 67.89; H, 4.62; N, 4.98.

trans-(4-Methoxyphenyl)(3-(2-nitrophenyl)oxiran-2-yl)methanone (6b).



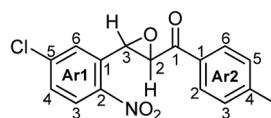
White powder (8.08 g, 0.027 mol, 100% yield): mp 141 $^{\circ}\text{C}$; ^1H NMR (400 MHz, DMSO- d_6) δ 8.21 (d, J = 8.2 Hz, 1H, H3-Ar1), 8.09 (d, J = 8.1 Hz, 2H, H2,6-Ar2), 7.87 (dd, J = 7.6, 7.4 Hz, 1H, H5-Ar1), 7.65–7.71 (m, 2H, H6,4-Ar1), 7.08 (d, J = 8.1 Hz, 2H, H3,5-Ar2), 4.71 (d, J = 1.7 Hz, 1H, H2), 4.56 (d, J = 1.7 Hz, 1H, H3), 3.87 (s, 3H, OMe); ^{13}C NMR (125.8 MHz, DMSO- d_6) δ 190.82 (1C=O), 163.87 (C4-Ar2), 147.45 (C2-Ar1), 134.60 (C5-Ar1), 132.53 (C1-Ar1), 130.87 (C2-Ar2), 129.43 (C4-Ar1), 128.20 (C1-Ar2), 127.05 (C6-Ar1), 124.61 (C3-Ar1), 114.11 (C3-Ar2), 58.17 (C2), 57.03 (C3), 55.58 (OMe). IR (nujol): ν 1678, 1597, 1513; HRMS (MALDI) calcd for $\text{C}_{16}\text{H}_{13}\text{NO}_5$ [M + Cs]⁺ 431.9843, found 431.9813; anal. calcd for $\text{C}_{16}\text{H}_{13}\text{NO}_5$: C, 64.21; H, 4.38; N, 4.68; found: C, 64.39; H, 4.51; N, 4.77.

trans-(3-Nitrophenyl)(3-(2-nitrophenyl)oxiran-2-yl)methanone (6c).



Grey powder (8.49 g, 0.027 mol, 100% yield): mp 156–157 $^{\circ}\text{C}$; ^1H NMR (400 MHz, DMSO- d_6) δ 8.79 (s, 1H, H2-Ar2), 8.53 (d, J = 8.2 Hz, 2H, H4,6-Ar2), 8.23 (d, J = 8.0 Hz, 1H, H3-Ar1), 7.86–7.11 (m, 2H, H5-Ar1, H5-Ar2), 7.67–7.72 (m, 2H, H4,6-Ar1), 4.90 (d, J = 1.9 Hz, 1H, H2), 4.65 (d, J = 1.9 Hz, 1H, H3); ^{13}C NMR (125.8 MHz, DMSO- d_6) δ 191.77 (1C=O), 148.05 (C3-Ar2), 147.45 (C2-Ar1), 136.05 (C1-Ar2), 134.70 (C5-Ar1), 134.63 (C6-Ar2), 132.15 (C1-Ar1), 130.72 (C5-Ar2), 129.64 (C4-Ar1), 128.17 (C4-Ar2), 127.07 (C6-Ar1), 124.68 (C3-Ar1), 122.71 (C2-Ar2), 58.71 (C2), 57.73 (C3). IR (nujol): ν 1697, 1522 cm^{-1} ; HRMS (MALDI) calcd for $\text{C}_{15}\text{H}_{10}\text{N}_2\text{O}_6$ [M + Cs]⁺ 446.9588, found 446.9581; anal. calcd for $\text{C}_{15}\text{H}_{10}\text{N}_2\text{O}_6$: C, 57.28; H, 3.32; N, 8.78; found: C, 57.33; H, 3.21; N, 8.91.

trans-(3-(5-Chloro-2-nitrophenyl)oxiran-2-yl)(4-tolyl)methanone (6d).

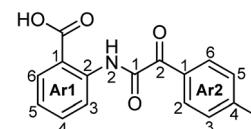


Tan powder (8.58 g, 0.027 mol, 100% yield): mp 139 $^{\circ}\text{C}$; ^1H NMR (400 MHz, DMSO- d_6) δ 8.24 (d, J = 8.8 Hz, 1H, H3-Ar1), 8.01 (d, J = 8.1 Hz, 2H, H2,6-Ar2), 7.75 (dd, J = 8.8, 2.4 Hz, 1H, H4-Ar1), 7.59 (d, J = 2.4 Hz, 1H, H6-Ar1), 7.37 (d, J = 8.1 Hz, 2H, H3,5-Ar2), 4.81 (d, J = 2.1 Hz, 1H, H2), 4.60 (d, J = 2.1 Hz, 1H, H3), 2.41 (s, 3H, Me); ^{13}C NMR (125.8 MHz, DMSO- d_6) δ 191.86 (1C=O), 146.17 (C2-Ar1), 144.85 (C4-Ar2), 139.44 (C5-Ar1), 134.86 (C1-Ar1), 132.64 (C1-Ar2), 129.41 (C3-Ar2), 129.35 (C4-Ar1), 128.56 (C2-Ar2), 126.82 (C3-Ar1), 126.71 (C6-Ar1), 58.24 (C2), 56.65 (C3), 21.20 (Me). IR (nujol): ν 1681, 1607, 1516; HRMS (MALDI) calcd for $\text{C}_{16}\text{H}_{12}\text{ClNO}_4$ [M + Cs]⁺ 449.9504, found 449.9522; anal. calcd for $\text{C}_{16}\text{H}_{12}\text{ClNO}_4$: C, 60.48; H, 3.81; Cl, 11.16; N, 4.41; found: C, 60.31; H, 3.71; N, 4.61.

Synthesis of 2-(2-oxo-2-arylacetamido)benzoic acids (7a–d).

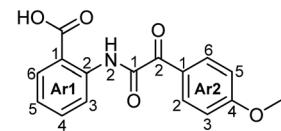
Synthesis of 2-(2-oxo-2-arylacetamido)benzoic acids (7a–d) was performed according to the general procedure for the rearrangement with the use of corresponding (3-(2-nitroaryl)oxiran-2-yl)(aryl)methanones (6a–d) instead of 3-(2-nitroaryl)oxiran-2-carboxamides (2a–h). Products 7a,b were purified by washing with acetone (3 \times 1 mL), products 7c,d – by recrystallization from AcOH.

2-[2-Oxo-2-(4-tolyl)acetamido]benzoic acid (7a).



Brown powder (0.35 g, 1.25 mmol, 75% yield): mp 199–201 $^{\circ}\text{C}$; ^1H NMR (400 MHz, DMSO- d_6) δ 12.72 (s, 1H, NH), 8.64 (d, J = 7.5 Hz, 1H, H3-Ar1), 8.14 (d, J = 7.8 Hz, 2H, H2,6-Ar2), 8.04 (dd, J = 8.1, 1.0 Hz, 1H, H6-Ar1), 7.71 (dd, J = 7.6, 7.5 Hz, 1H, H4-Ar1), 7.37 (d, J = 7.8 Hz, 2H, H3,5-Ar2), 7.28 (dd, J = 7.5, 7.9 Hz, 1H, H5-Ar1), 2.42 (s, 3H, Me); ^{13}C NMR (125.8 MHz, DMSO- d_6) δ 187.00 (2C=O), 169.15 (CO₂H), 160.57 (1C=O), 145.42 (C4-Ar2), 139.28 (C2-Ar1), 134.19 (C4-Ar1), 131.41 (C6-Ar1), 130.94 (C2-Ar2), 130.38 (C1-Ar2), 129.27 (C3-Ar2), 124.00 (C5-Ar1), 120.13 (C3-Ar1), 117.75 (C1-Ar1), 21.39 (Me). IR (nujol): ν 3363, 1678, 1604, 1586, 1521, 1262 cm^{-1} ; HRMS (MALDI) calcd for $\text{C}_{16}\text{H}_{13}\text{NO}_4$ [M + Cs]⁺ 415.9894, found 415.9887; anal. calcd for $\text{C}_{16}\text{H}_{13}\text{NO}_4$: C, 67.84; H, 4.63; N, 4.94. Found: C, 67.92; H, 4.68; N, 4.71.

2-[2-(4-Methoxyphenyl)2-oxoacetamido]benzoic acid (7b).

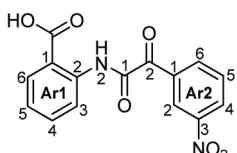


Brown powder (0.40 g, 1.35 mmol, 81% yield): mp 178–181 $^{\circ}\text{C}$; ^1H NMR (400 MHz, DMSO- d_6) δ 12.37 (s, 1H, NH), 8.65 (d, J = 8.3 Hz, 1H, H3-Ar1), 8.27 (d, J = 8.9 Hz, 2H, H2,6-Ar2), 8.05 (d, J = 8.0 Hz, 1H, H6-Ar1), 7.67 (dd, J = 7.2, 8.3 Hz, 1H, H4-Ar1), 7.26 (dd, J = 7.2, 7.8 Hz, 1H, H5-Ar1), 8.10 (d, J = 8.9 Hz, 2H, H3,5-Ar2), 3.88 (s, 3H, OMe); ^{13}C NMR (125.8 MHz, DMSO- d_6) δ 185.42 (1C=O), 169.12 (CO₂H), 164.39 (C4-Ar2), 160.81 (2C=O), 139.35 (C2-Ar1), 134.14 (C4-Ar1), 133.47 (C2-Ar2), 131.39 (C6-



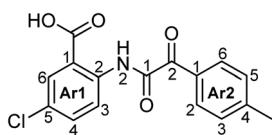
Ar1), 125.61 (C4-Ar2), 123.88 (C5-Ar1), 120.04 (C3-Ar1), 117.60 (C1-Ar1), 114.10 (C3-Ar2), 55.70 (OMe). IR (nujol): ν 3472, 3258, 1702, 1674, 1601, 1584, 1528, 1260 cm^{-1} ; HRMS (MALDI) calcd for $\text{C}_{16}\text{H}_{13}\text{NO}_5$ [M + Cs]⁺ 431.9843, found 431.9828; anal. calcd for $\text{C}_{16}\text{H}_{13}\text{NO}_5$: C, 64.21; H, 4.38; N, 4.68. Found: C, 64.32; H, 4.37; N, 4.75.

2-[2-(3-Nitrophenyl)-2-oxoacetamido]benzoic acid (7c).



Brown powder (0.45 g, 1.44 mmol, 86% yield): mp 251–252 °C; ¹H NMR (400 MHz, DMSO-*d*₆) δ 12.60 (s, 1H, NH), 9.06 (s, 1H, H2-Ar2), 8.70 (d, *J* = 8.1 Hz, 1H, H3-Ar1), 8.58 (d, *J* = 7.7 Hz, 1H, H6-Ar2), 8.54 (dd, *J* = 8.2, 1.5 Hz, 1H, H4-Ar2), 8.09 (dd, *J* = 7.9, 1.5 Hz, 1H, H6-Ar1), 7.89 (dd, *J* = 7.9, 8.0 Hz, 1H, H5-Ar2), 7.72 (ddd, *J* = 7.9, 7.9, 1.5 Hz, 1H, H4-Ar1), 7.30 (dd, *J* = 7.9, 7.9 Hz, 1H, H5-Ar1); ¹³C NMR (125.8 MHz, DMSO-*d*₆) δ 185.04 (1C=O), 169.14 (CO₂H), 159.14 (2C=O), 147.38 (C3-Ar2), 139.26 (C2-Ar1), 136.71 (C6-Ar2), 134.46 (C1-Ar2), 134.22 (C4-Ar1), 131.44 (C6-Ar1), 130.14 (C5-Ar2), 128.038 (C4-Ar2), 125.58 (C2-Ar2), 123.99 (C5-Ar1), 119.88 (C3-Ar1), 117.58 (C1-Ar1). IR (nujol): ν 3413, 3189, 1695, 1679, 1588, 1532, 1288 cm^{-1} ; HRMS (MALDI) calcd for $\text{C}_{15}\text{H}_{10}\text{N}_2\text{O}_6$ [M + 2Cs – H]⁺ 578.8564, found 578.8545; anal. calcd for $\text{C}_{15}\text{H}_{10}\text{N}_2\text{O}_6$: C, 57.28; H, 3.32; N, 8.78; found: C, 57.34; H, 3.37; N, 8.89.

5-Chloro-2-[2-oxo-2-(4-tolyl)acetamido]benzoic acid (7d).



Light yellow powder (0.47 g, 1.49 mmol, 89% yield): mp 257–258 °C; ¹H NMR (400 MHz, DMSO-*d*₆) δ 12.30 (s, 1H, NH), 8.64 (d, *J* = 8.9 Hz, 1H, H3-Ar1), 8.13 (d, *J* = 7.9 Hz, 2H, H2,6-Ar2), 7.97 (br.s, 1H, H6-Ar1), 7.74 (dd, *J* = 8.8, 1.8 Hz, 1H, H4-Ar1), 7.38 (d, *J* = 7.9 Hz, 2H, H3,5-Ar2), 2.41 (s, 3H, Me); ¹³C NMR (125.8 MHz, DMSO-*d*₆) δ 186.46 (1C=O), 167.80 (CO₂H), 160.35 (2C=O), 145.35 (C4-Ar2), 138.03 (C2-Ar1), 133.70 (C4-Ar1), 130.90 (C2-Ar2), 130.51 (C6-Ar1), 130.25 (C1-Ar2), 129.16 (C3-Ar2), 127.55 (C5-Ar1), 121.83 (C3-Ar1), 119.53 (C1-Ar1), 21.32 (Me). IR (nujol): ν 3219, 1703, 1681, 1598, 1577, 1521, 1248 cm^{-1} ; HRMS (MALDI) calcd for $\text{C}_{16}\text{H}_{12}\text{ClNO}_4$ [M + 2Cs – H]⁺ 581.8480, found 581.8455; anal. calcd for $\text{C}_{16}\text{H}_{12}\text{ClNO}_4$: C, 60.48; H, 3.81; Cl, 11.16; N, 4.41; found: C, 60.52; H, 3.73; N, 4.58.

Conflict of interest

The authors declare no competing financial interest.

Acknowledgements

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