Masashige Yamauchi

Faculty of Pharmaceutical Sciences, Josai University, Keyakidai, Sakado, Saitama 350-0295, Japan Received May 13, 2002

The reaction of 2-methylene-1,3-dicarbonyl compounds 1 and nitrile oxides, which were prepared from hydroxymoyl chlorides 2 with triethylamine, gave 5,5-disubstituted 2-isoxazolines 3 regioselectively.

J. Heterocyclic Chem., 39, 1013 (2002).

The formation of 2-isoxazolines by 1,3-dipolar cycloaddition of nitrile oxides to alkenes has been known for many years, and the scope and mechanism of the reaction have been studied in detail [1]. The utility of this heterocycle is derived from its ready conversion to useful synthetic intermediates, β -hydroxy ketones[2] and γ -amino alcohols [3]. In general, with monosubstituted olefins reaction gives exclusively or predominantly the 5-substituted isoxazoline, whatever the nature of the substituent on the dipolarophile. Reactions of nitrile oxide with both electron-rich and electron-deficient dipolarophiles are controlled mainly by the LUMO (dipole) - HOMO (dipolarophile) interaction, and union of the atoms with the larger coefficients leads to 5substituted isoxazolines [4]. Houk et al. predicted that unsymmetrical electron-deficient 1,1-disubstituted ethylene will react with nitrile oxide to produce predominantly 4,4-disubstituted 2-isoxazolines [4b]. During the course of our studies on the reactivity of 2-methylene-1,3-dicarbonyl compounds 1 having two electron-withdrawing groups at the same olefinic carbon [5] we investigated the reaction with nitrile oxide. It is expected that the introduction of carbonyl group at the α -position of the α , β -unsaturated carbonyl compounds lower the LUMO energy level therefore its reaction with nitrile oxide is controlled by LUMO (dipolarophile) - HOMO (dipole) interaction. The coefficients of the terminal methylene carbons obtained from MNDO calculation of the most stable conformer of ethyl 2-benzoylacrylate 1d are shown in Figure 1. Taking this into consideration with the coefficients of benzonitrile oxide reported by Houk et al. [4] the addition product could be 4,4-disub-





stituted 2-isoxazoline **4d-a** whether the reaction is controlled by LUMO (dipolarophile) (-0.6 eV) - HOMO(dipole) (-10 eV) or HOMO (dipolarophile) (-11 eV) -LUMO (dipole) (-1.0 eV) interactions.

Triethylamine was added to a solution of enedione 1, hydroxymoyl chloride 2, which was prepared by Larsen's procedure [6], in CH_2Cl_2 . The color of the reaction mixture turned pale yellow immediately. The solvent was evaporated off and the resulting residue was applied to preparative thin layer chromatography to give the product. Other cycloaddition products were not detected.

The C-4 methylene protons of the product each appeared as a doublet at *ca*. 3.7 ppm and *ca*. 4.5 ppm except for dibenzoyl **3b** series in the ¹H nmr spectra. From these results, it is not possible to distinguish whether the product is 5,5-disubstituted 2-isoxazoline **3** or 4,4-disubstituted 2isoxazoline **4** clearly. The methylene carbons of the product show the signals at *ca*. 41~45 ppm in the ¹³C nmr spectra. The ring methylene carbon of pyrazoline **5** resonance at 39.2 ppm [5d] whereas that of isoxazolizine **6** appear at 72.1 ppm [7], which suggest the product must be 5,5-disubstituted 2-isoxazoline **3**. This is further supported by observing the correlations in the HMBC (heteronuclear multiple bond correlation) spectrum of **3d-a** between signals of C-4 methylene protons and C-3, C-5, both carbonyl carbons and quarternary phenyl carbon attached to C-3.



The dipolar addition products were 5,5-disubstituted 2-isoxazolines **3** not 4,4-disubstituted 2-isoxazolines **4** predicted from frontier orbital. Huisgen *et al.* reported that

Table 1 4,5-Dihydroisoxazoles **3**

| Entry | \mathbb{R}^1 | R ² | Yield(%)[a] |
|-------|----------------|-----------------------|-------------|
| a-a | Me | Н | 66 |
| a-b | Me | Me | 76 |
| a-c | Me | Pr ⁱ | 62 |
| a-d | Me | OMe | 72 |
| a-f | Me | Br | 67 |
| a-g | Me | naphthyl | 67 |
| b-a | Ph | Н | 72 |
| b-b | Ph | Me | 82 |
| b-c | Ph | Pr ⁱ | 70 |
| b-d | Ph | OMe | 70 |
| b-e | Ph | Cl | 70 |
| b-f | Ph | Br | 73 |
| b-g | Ph | naphthyl | 74 |
| c-a | OMe | Н | 94 |
| c-b | OMe | Me | 84 |
| c-c | OMe | Pr ⁱ | 71 |
| c-e | OMe | Cl | 82 |
| c-f | OMe | Br | 89 |
| c-g | OMe | naphthyl | 96 |
| d-a | OEt | Н | 75 |
| d-b | OEt | Me | 84 |
| d-c | OEt | Pr ⁱ | 72 |
| d-d | OEt | OMe | 72 |
| d-e | OEt | Cl | 85 |
| d-f | OEt | Br | 80 |
| d-g | OEt | naphthyl | 76 |
| e-a | O-cyclohexyl | Н | 95 |
| e-b | O-cyclohexyl | Me | 99 |
| e-c | O-cyclohexyl | Pr ⁱ | 98 |
| e-d | O-cyclohexyl | OMe | 77 |
| e-e | O-cyclohexyl | Cl | 88 |
| e-f | O-cyclohexyl | Br | 92 |
| e-g | O-cyclohexyl | naphthyl | 98 |
| f-a | O-benzyl | Н | 70 |
| f-b | O-benzyl | Me | 88 |
| f-c | O-benzyl | Pr ⁱ | 83 |
| f-d | O-benzyl | OMe | 71 |
| f-e | O-benzyl | Cl | 89 |
| f-f | O-benzyl | Br | 90 |
| f-g | O-benzyl | naphthyl | 82 |

[a] Isolated yields.



selected HMBC correlations of 3d-a

nitrile oxides reacted with methyl acrylate to give 2-isoxazoline-5-carboxylic esters predominantly whereas almost equal ratio of 4- and 5-carboxylates were formed with methyl crotonate and methyl cinnamate [8]. This means that not only frontier orbital factor but also steric factor controls the reaction. In the reaction of 1,1-disubstituted ethylene such as 2-methylene-1,3-dicarbonyl compound with nitrile oxide, steric hindrance factor overcomes frontier orbital factor resulting in the formation of 5,5-disubstituted 2-isoxazoline.

EXPERIMENTAL

The nmr spectra were measured on a JEOL GX-270 spectrometer for samples in deuteriochloroform solution at 270 MHz for ¹H and 67.89 MHz for ¹³C, and chemical shifts are expressed in δ -units using tetramethylsilane or chloroform as an internal reference. The ir spectra of solids (potassium bromide) and liquids (film) were recorded on a JASCO FT/IR-410 spectrometer. Highresolution mass spectra were obtained with a JEOL JMS-700 spectrometer. To dry the organic layers after extraction MgSO₄ was used. Preparative thin layer chromatography was performed with Silica gel 60 F₂₅₄ 0.5 mm (Merck).

General Procedure for the Preparation of 4,5-Dihydroisoxazoles 3.

To a stirred solution of 2-methylene-1,3-dicarbonyl compounds (1, 0.5 mmol) and hydroxymoyl chlorides (2, 0.5 mmol) in CH_2Cl_2 (3 ml), triethylamine (70 µl, 0.05 mmol) was added in one portion with stirring. The reaction mixture turned pale yellow immediately. After stirring was continued for additional 5 minutes, the reaction mixture was washed with water (3 ml x 5) and brine (2 ml). The organic layer was dried and evaporated. The resulting residue was subjected to preparative thin layer chromatography to give **3**. The yields of **3** are shown in table 1.

5-Acetyl-5-benzoyl-3-phenyl-4,5-dihydroisoxazole (3a-a).

This compound was obtained as viscous oil; ir v CO 1717, 1688 cm⁻¹; ¹H nmr: δ 2.45 (s, 3H, CH₃), 3.82 (d, J = 17.8 Hz, 1H, CHH), 4.28 (d, J = 17.8 Hz, 1H, CHH), 7.42-7.73 (m, 8H, ArH), 8.01 (br d, J = 7.1 Hz, 2H, ArH); ¹³C nmr: δ 25.5, 41.4, 97.3, 127.0, 127.8, 128.7, 128.8, 129.8, 130.9, 133.2, 134.0, 156.4, 192.3, 202.9; HR-MS m/z: Calcd. for C₁₈H₁₅NO₃: 293.1052. Found: 293.1060.

5-Acetyl-5-benzoyl-3-(4-methylphenyl)-4,5-dihydroisoxazole (**3a-b**).

This compound was obtained as viscous oil; ir v CO 1719, 1687 cm^{-1.} ¹H nmr: δ 2.39 (s, 3H, CH₃), 2.44 (s, 3H, CH₃), 3.81 (d, J = 17.6 Hz, 1H, CHH), 4.26 (d, J = 17.6 Hz, 1H, CHH), 7.23 (d, J = 8.1 Hz, 2H, ArH), 7.47 (br t, J = 7.3 Hz, 2H, ArH), 7.59 (d, J = 8.1 Hz, 2H, ArH), 7.60 (br t, J = 7.3 Hz, 1H, ArH), 8.01 (br d, J = 7.3 Hz, 2H, ArH); ¹³C nmr: δ 21.6, 25.5, 41.5, 97.1, 124.9, 126.9, 128.6, 129.5, 129.8, 133.2, 133.9, 141.3, 156.3, 192.5, 203.0; HR-MS m/z: Calcd. for C₁₉H₁₇NO₃: 307.1208. Found: 307.1225.

5-Acetyl-5-benzoyl-3-(4-isopropylphenyl)-4,5-dihydroisoxazole (**3a-c**).

This compound was obtained as viscous oil; ir v CO 1719, 1690 cm⁻¹. ¹H nmr: δ 1.26 (d, J = 6.9 Hz, 6H, CH₃), 2.44 (s, 3H,

CH₃), 2.94 (heptet, J = 6.9 Hz, 1H, CH), 3.81 (d, J = 17.6 Hz, 1H, CHH), 4.26 (d, J = 17.6 Hz, 1H, CHH), 7.29 (d, J = 8.3 Hz, 2H, ArH), 7.47 (br t, J = 7.3Hz, 2H, ArH), 7.59 (br t, J = 7.3 Hz, 1H, ArH), 7.63 (d, J = 8.3 Hz, 2H, ArH), 8.00 (br d, J = 7.3 Hz, 2H, ArH); ¹³C nmr: δ 23.8, 25.4, 34.2, 41.6, 97.1, 125.3, 126.9, 127.1, 128.6, 129.8, 133.2, 133.9, 152.1, 156.3, 192.5, 203.1; HR-MS m/z: Calcd. for C₂₁H₂₁NO₃: 335.1521. Found: 335.1509.

5-Acetyl-5-benzoyl-3-(4-methoxyphenyl)-4,5-dihydroisoxazole (**3a-d**).

This compound was obtained as colorless needles (hexane- C_6H_6), mp 111-112°, ir v CO 1715, 1689 cm⁻¹; ¹H nmr: δ 2.44 (s, 3H, CH₃), 3.79 (d, J = 17.6 Hz, 1H, CHH), 3.85 (s, 3H, CH₃), 4.25 (d, J = 17.6 Hz, 1H, CHH), 6.93 (d, J = 9.0 Hz, 2H, ArH), 7.47 (br t, J = 9.0 Hz, 2H, ArH), 7.57 (br t, J = 7.3 Hz, 1H, ArH), 7.64 (d, J = 9.0 Hz, 2H, ArH), 8.01 (br d, J = 7.3 Hz, 2H, ArH); ¹³C nmr: δ 25.5, 41.6, 55.4, 97.1, 114.2, 120.3, 128.59, 128.63, 129.8, 133.2, 133.9, 155.9, 161.5, 192.6, 203.2; HR-MS m/z: Calcd. for C₁₉H₁₇NO₄: 323.1158. Found: 323.1154.

5-Acetyl-5-benzoyl-3-(4-bromophenyl)-4,5-dihydroisoxazole (**3a-f**).

This compound was obtained as viscous oil; ir v CO 1717, 1687 cm⁻¹. ¹H nmr: δ 2.44 (s, 3H, CH₃), 3.79 (d, J = 17.5 Hz, 1H, CHH), 4.25 (d, J = 17.5 Hz, 1H, CHH), 7.47-7.61 (m, 7H, ArH), 7.99 (br d, J = 7.1 Hz, 2H, ArH); ¹³C nmr: δ 25.5, 41.1, 97.6, 125.3, 126.7, 128.4, 128.7, 129.8, 132.1, 133.1, 134.1, 155.6, 192.1, 202.5; HR-MS m/z: Calcd. for C₁₈H₁₄BrNO₃: 371.0157. Found: 371.0175.

5-Acetyl-5-benzoyl-3-naphthalen-1-yl-4,5-dihydroisoxazole (**3a-g**).

This compound was obtained as viscous oil; ir v CO 1718, 1687 cm⁻¹. ¹H nmr: δ 2.50 (s, 3H, CH₃), 4.02 (d, J = 17.5 Hz, 1H, CHH), 4.50 (d, J = 17.5 Hz, 1H, CHH), 7.46-7.96 (m, 9H, ArH), 8.06 (br d, J = 7.1 Hz, 2H, ArH), 8.96 (br d, J = 8.9 Hz, 1H, ArH); ¹³C nmr: δ 25.5, 44.1, 96.1, 124.7, 126.5, 126.7, 127.8, 128.2, 128.5, 128.6, 128.7, 129.8, 130.3, 131.2, 133.3, 133.9, 134.0, 157.0, 192.4, 203.0; HR-MS m/z: Calcd. for C₂₂H₁₇NO₃: 343.1208. Found: 343.1212.

5,5-Dibenzoyl-3-phenyl-4,5-dihydroisoxazole (3b-a).

This compound was obtained as colorless needles (hexane- C_6H_6), mp 105-106°, ir v CO 1699, 1680 cm⁻¹; ¹H nmr: δ 4.32 (s, 2H, CH₂), 7.40-7.77 (m, 11H, ArH), 8.03 (br d, J = 7.2 Hz, 4H, ArH); ¹³C nmr: δ 42.3, 96.7, 127.1, 127.9, 128.7, 128.8, 130.0, 130.8, 133.0, 133.9, 156.5, 192.7.

Anal. Calcd. for C₂₃H₁₇NO₃: C, 77.73; H, 4.82; N, 3.84. Found: C, 77.58; H, 4.78; N, 3.74.

5,5-Dibenzoyl-3-(4-methylphenyl)-4,5-dihydroisoxazole (3b-b).

This compound was obtained as colorless needles (hexane- C_6H_6), mp 125-127°, ir v CO 1700, 1669 cm⁻¹; ¹H nmr: δ 2.39 (s, 3H, CH₃), 4.30 (s, 2H, CH₂), 7.43 (br t, J = 7.5 Hz, 4H, ArH), 7.56 (br t, J = 7.5 Hz, 2H, ArH), 7.63 (d, J = 8.2 Hz, 2H, ArH), 8.03 (br d, J = 7.5 Hz, 4H, ArH); ¹³C nmr: δ 21.5, 42.3, 96.6, 125.2, 127.1, 128.7, 129.6, 130.1, 133.2, 134.0, 141.3, 156.6, 193.1.

Anal. Calcd. for $C_{24}H_{19}NO_3$: C, 78.03; H, 5.18; N, 3.79. Found: C, 78.23; H, 5.07; N, 3.66.

5,5-Dibenzoyl-3(4-isopropylphenyl)-4,5-dihydroisoxazole (**3b-c**).

This compound was obtained as colorless needles (hexane- C_6H_6), mp 102-103°, ir v CO 1691, 1671 cm⁻¹; ¹H nmr: δ 1.26 (d, J = 6.9 Hz, 6H, CH₃), 2.96 (heptet, J = 6.9 Hz, 1H, CH), 4.30 (s, 2H, CH₂), 7.29 (d, J = 8.2 Hz, 2H, ArH), 7.43 (br t, J = 7.4 Hz, 4H, ArH), 7.56 (br t, J = 7.4 Hz, 2H, ArH), 7.67 (d, J = 8.2 Hz, 2H, ArH), 8.03 (br d, J = 7.4 Hz, 4H, ArH); ¹³C nmr: δ 23.8, 34.2, 42.4, 96.6, 125.5, 126.9, 127.2, 128.6, 130.0, 133.1, 133.9, 152.1, 156.4, 192.9.

Anal. Calcd. for $C_{26}H_{23}NO_3$: C, 78.57; H, 5.83; N, 3.52. Found: C, 78.45; H, 5.77; N, 3.36.

5,5-Dibenzoyl-3-(4-methoxyphenyl)-4,5-dihydroisoxazole (3b-d).

This compound was obtained as viscous oil; ir v CO 1696, 1679 cm⁻¹. ¹H nmr: δ 3.84 (s, 3H, CH₃), 4.29 (s, 2H, CH₂), 6.93 (d, J = 9.0 Hz, 2H, ArH), 7.43 (br t, J = 7.1 Hz, 4H, ArH), 7.56 (br t, J = 7.1 Hz, 2H, ArH), 7.67 (d, J = 9.0 Hz, 2H, ArH), 8.03 (br d, J = 7.1 Hz, 4H, ArH); ¹³C nmr: δ 42.5, 55.4, 96.5, 114.2, 120.4, 128.6, 128.7, 130.0, 133.1, 133.8, 156.0, 161.5, 193.0; HR-MS m/z: Calcd. for C₂₄H₁₉NO₄: 385.1314. Found: 385.1344.

5,5-Dibenzoyl-3-(4-chlorophenyl)-4,5-dihydroisoxazole (3b-e).

This compound was obtained as colorless needles (hexane- C_6H_6), mp 152-153°, ir v CO 1699, 1666 cm⁻¹; ¹H nmr: δ 4.28 (s, 2H, CH₂), 7.35-7.69 (m, 10H, ArH), 8.02 (br d, J = 7.2 Hz, 4H, ArH); ¹³C nmr: δ 42.0, 97.0, 126.5, 128.3, 128.7, 129.1, 130.0, 133.0, 134.0, 136.9, 155.6, 192.5.

Anal. Calcd. for C₂₃H₁₆ClNO₃: C, 70.86; H, 4.14; N, 3.59. Found: C, 71.10; H, 4.06; N, 3.40.

5,5-Dibenzoyl-3-(4-bromophenyl)-4,5-dihydroisoxazole (3b-f).

This compound was obtained as colorless needles (hexane- C_6H_6), mp 161-162°, ir v CO 1699, 1667 cm⁻¹; ¹H nmr: δ 4.28 (s, 2H, CH₂), 7.35-7.63 (m, 7H, ArH), 8.01 (br d, J = 7.4 Hz, 2H, ArH); ¹³C nmr: δ 41.9, 97.0, 125.5, 126.9, 128.4, 128.7, 130.0, 132.0, 132.9, 134.0, 155.7, 192.4; HR-MS m/z: Calcd. for C₁₉H₁₆BrNO₄: 433.0314. Found: 433.0300.

5,5-Dibenzoyl-3-naphthalen-1-yl-4,5-dihydroisoxazole (**3b-g**).

This compound was obtained as colorless needles (hexane- C_6H_6), mp 154-155°, ir v CO 1698, 1670 cm⁻¹; ¹H nmr: δ 4.52 (s, 2H, CH₂), 7.41-7.96 (m, 12H, ArH), 8.09 (br d, J = 7.1 Hz, 4H, ArH), 8.99 (br d, J = 8.2 Hz, 1H, ArH); ¹³C nmr: δ 45.0, 95.6, 124.7, 124.9, 126.5, 126.9, 127.8, 128.5, 128.6, 128.7, 130.0, 130.4, 131.6, 133.1, 133.86, 133.93, 157.1, 192.8.

Anal. Calcd. for $C_{25}H_{19}NO_3$: C, 79.98; H, 4.72; N, 3.45. Found: C, 80.06; H, 4.69; N, 3.41.

5-Benzoyl-3-phenyl-4,5-dihydroisoxazole-5-carboxylic acid methyl ester. (**3c-a**).

This compound was obtained as colorless needles (hexane- C_6H_6), mp 134-135°, ir v CO 1734, 1696 cm⁻¹; ¹H nmr: δ 3.74 (d, J = 17.6 Hz, 1H, CHH), 3.79 (3H, s, CH₃), 4.55 (d, J = 17.6 Hz, 1H, CHH), 7.35-7.72 (m, 8H, ArH), 8.12 (br d, J = 7.1 Hz, 2H, ArH); ¹³C nmr: δ 42.4, 53.6, 92.0, 126.9, 127.9, 128.6, 128.7, 130.0, 130.7, 133.1, 134.0, 156.1, 169.5, 189.8.

Anal. Calcd. for C₁₈H₁₅NO₄: C, 69.89; H, 4.89; N, 4.53. Found: C, 69.96; H, 4.86; N, 4.53.

5-Benzoyl-3-(4-methylphenyl)-4,5-dihydroisoxazole-5-carboxylic Acid Methyl Ester (**3c-b**).

This compound was obtained as colorless needles (hexane- C_6H_6), mp 99°, ir v CO 1754, 1691 cm⁻¹; ¹H nmr: δ 3.73 (d, J = 17.6 Hz, 1H, *CH*H), 4.39 (s, 3H, CH₃), 4.53 (d, J = 17.6 Hz, 1H, CH*H*), 7.20-7.60 (m, 7H, ArH), 8.12 (br d, J = 7.3 Hz, 2H, ArH); ¹³C nmr: δ 21.5, 53.6, 77.2, 91.9, 125.2, 127.0 128.7, 129.5, 130.1, 133.3, 134.0, 141.2, 156.2, 169.2, 190.2.

Anal. Calcd. for $C_{19}H_{17}NO_4$: C, 70.58; H, 5.30; N, 4.33. Found: C, 70.55; H, 5.11; N, 4.22.

5-Benzoyl-3-(4-isopropylphenyl)-4,5-dihydroisoxazole-5-carboxylic Acid Methyl Ester (**3c-c**).

This compound was obtained as colorless needles (hexane), mp 67°, ir v CO 1728, 1697 cm⁻¹; ¹H nmr: δ 1.26 (d, J = 6.9 Hz, 6H, CH₃), 2.94 (heptet, J = 6.9 Hz, 1H, CH), 3.73 (d, J = 17.6 Hz, 1H, CHH), 3.78 (s, 3H, CH₃), 4.52 (d, J = 17.6 Hz, 1H, CHH), 7.25-7.65 (m, 7H, ArH), 8.12 (br d, J = 7.2 Hz, 2H, ArH); ¹³C nmr: δ 23.8, 34.2, 42.6, 53.6, 91.9, 125.4, 126.8, 127.0, 128.6, 130.0, 133.2, 133.9, 152.0, 156.0, 169.5, 190.0.

Anal. Calcd. for C₂₁H₂₁NO₄: C, 71.78; H, 6.02; N, 3.99. Found: C, 71.72; H, 5.99; N, 3.97.

5-Benzoyl-3-(4-chlorophenyl)-4,5-dihydroisoxazole-5-carboxylic Acid Methyl Ester (**3c-d**).

This compound was obtained as colorless needles (hexane- C_6H_6), mp 121-122°, ir v CO 1752, 1684 cm⁻¹; ¹H nmr: δ 3.70 (d, J = 17.6 Hz, 1H, CHH), 3.79 (s, 3H, CH₃), 4.53 (d, J = 17.6 Hz, 1H, CHH), 7.25-7.65 (m, 7H, ArH), 8.11 (br d, J = 7.1 Hz, 2H, ArH); ¹³C nmr: δ 42.2, 53.7, 92.2, 126.4, 128.2, 128.6, 129.0, 130.0, 133.0, 134.1, 136.8, 155.2, 169.3, 189.6.

Anal. Calcd. for $C_{18}H_{14}CINO_4$: C, 62.89; H, 4.10; N, 4.07. Found: C, 62.96; H, 4.01; N, 4.08.

5-Benzoyl-3-(4-bromophenyl)-4,5-dihydroisoxazole-5-carboxylic Acid Methyl Ester (**3c-f**).

This compound was obtained as colorless needles (hexane- C_6H_6), mp 130-131°, ir v CO 1755, 1691 cm⁻¹; ¹H nmr: δ 3.70 (d, J = 17.6 Hz, 1H, CHH), 3.79 (s, 3H, CH₃), 4.53 (d, J = 17.6 Hz, 1H, CHH), 7.25-7.65 (m, 7H, ArH), 8.11 (br d, J = 7.1 Hz, 2H, ArH); ¹³C nmr: δ 42.1, 53.7, 92.2, 125.2, 126.8, 128.3, 128.6, 130.0, 132.0, 133.0, 134.1, 155.3, 169.3, 189.5.

Anal. Calcd. for $C_{18}H_{14}BrNO_4$: C, 55.69; H, 3.63; N, 3.61. Found: C, 55.86; H, 3.51; N, 3.62.

5-Benzoyl-3-naphthalen-1-yl-4,5-dihydroisoxazole-5-carboxylic acid methyl ester (**3c-g**).

This compound was obtained as colorless needles (hexane- C_6H_6), mp 130-132°, ir v CO 1761, 1696 cm⁻¹; ¹H nmr: δ 3.81 (s, 3H, CH₃), 3.92 (d, J = 17.5 Hz, 1H, CHH), 4.78 (d, J = 17.5 Hz, 1H, CHH), 7.46-7.95 (m, 9H, ArH), 8.17 (br d, J = 7.1 Hz, 2H, ArH), 8.96 (br d, J = 8.2 Hz, 1H, ArH); ¹³C nmr: δ 45.1, 53.7, 90.8, 124.6, 124.8, 126.4, 126.9, 127.7, 128.3, 128.5, 128.6, 130.0, 130.4, 131.5, 133.8, 134.0, 156.7, 169.6, 189.8.

Anal. Calcd. for C₂₂H₁₇NO₄: C, 73.53; H, 4.77; N, 3.90. Found: C, 73.38; H, 4.63; N, 3.90.

5-Benzoyl-3-phenyl-4,5-dihydroisoxazole-5-carboxylic Acid Ethyl Ester (**3d-a**).

This compound was obtained as colorless needles (hexane- C_6H_6), mp 98-99°, ir v CO 1727, 1694 cm⁻¹; ¹H nmr: δ 1.16 (t,

 $\begin{array}{l} J=7.1 \ \text{Hz}, \ 3\text{H}, \ \text{CH}_3), \ 3.71 \ (\text{d}, \ J=17.5 \ \text{Hz}, \ 1\text{H}, \ \text{CH}\text{H}), \ 4.25 \ (\text{q}, \\ J=7.1 \ \text{Hz}, \ 2\text{H}, \ \text{OCH}_2), \ 4.56 \ (\text{d}, \ J=17.5 \ \text{Hz}, \ 1\text{H}, \ \text{CH}\text{H}), \ 7.39 \\ 7.73 \ (\text{m}, \ 8\text{H}, \ \text{ArH}), \ 8.13 \ (\text{br} \ \text{d}, \ J=7.0 \ \text{Hz}, \ 2\text{H}, \ \text{ArH}); \ ^{13}\text{C} \ \text{nmr}; \ \delta \\ 13.8, \ 42.2, \ 62.8, \ 92.0, \ 127.0, \ 128.1, \ 128.6, \ 128.8, \ 130.1, \ 130.8, \\ 133.3, \ 134.0, \ 156.1, \ 169.1, \ 190.0. \end{array}$

Anal. Calcd. for C₁₉H₁₇NO₄: C, 70.58; H, 5.30; N, 4.33. Found: C, 70.49; H, 5.30; N, 4.00.

5-Benzoyl-3-(4-methylphenyl)-4,5-dihydroisoxazole-5-carboxylic Acid Ethyl Ester (**3d-b**).

This compound was obtained as colorless needles (hexane- C_6H_6), mp 87°, ir v CO 1756, 1695 cm⁻¹; ¹H nmr: δ 1.16 (t, J = 7.1 Hz, 3H, CH₃), 2.38 (s, 3H, CH₃), 3.69 (d, J = 17.5 Hz, 1H, CHH), 4.24 (q, J = 7.1 Hz, 2H, OCH₂), 4.53 (d, J = 17.5 Hz, 1H, CHH), 7.20-7.61 (m, 7H, ArH), 8.13 (br d, J = 7.3 Hz, 2H, ArH); ¹³C nmr: δ 13.8, 21.5, 42.3, 62.8, 91.8, 125.2, 127.0, 128.6, 129.5, 130.1, 133.4, 133.9, 141.2, 156.1, 169.2, 190.2.

Anal. Calcd. for C₂₀H₁₉NO₄: C, 71.20; H, 5.68; N, 4.15. Found: C, 71.08; H, 5.50; N, 4.03.

5-Benzoyl-3-(4-isopropylphenyl)-4,5-dihydroisoxazole-5-carboxylic Acid Ethyl Ester (**3d-c**).

This compound was obtained as colorless needles (hexane), mp 77-80°, ir v CO 1753, 1693 cm⁻¹; ¹H nmr: δ 1.16 (t, J = 7.1 Hz, 3H, CH₃), 1.26 (d, J = 6.9 Hz, 6H, CH₃), 2.93 (heptet, J = 6.9 Hz, 1H, CH), 3.70 (d, J = 17.5 Hz, 1H, CHH), 4.24 (q, J = 7.1 Hz, 2H, OCH₂), 4.53 (d, J = 17.5 Hz, 1H, CHH), 7.25-7.65 (m, 7H, ArH), 8.12 (br d, J = 7.1 Hz, 2H, ArH); ¹³C nmr: δ 13.9, 23.8, 34.2, 42.4, 62.8, 91.8, 125.5, 126.8, 127.0, 128.5, 130.0, 133.3, 133.8, 151.9, 155.9, 169.0, 190.0.

Anal. Calcd. for C₂₂H₂₃NO₄: C, 72.31; H, 6.34; N, 3.83. Found: C, 72.13; H, 6.27; N, 3.54.

5-Benzoyl-3-(4-methoxyphenyl)-4,5-dihydroisoxazole-5-carboxylic Acid Ethyl Ester (**3d-d**).

This compound was obtained as colorless needles (hexane- C_6H_6), mp 86-87°, ir v CO 1755, 1695 cm⁻¹; ¹H nmr: δ 1.16 (t, J = 7.1 Hz, 3H, CH₃), 3.68 (d, J = 17.5 Hz, 1H, CHH), 3.84 (s, 3H, CH₃), 4.24 (q, J = 7.1 Hz, 2H, OCH₂), 4.52 (d, J = 17.5 Hz, 1H, CHH), 6.92 (d, J = 8.8 Hz, 2H, ArH), 7.47 (br t, J = 7.3 Hz, 2H, ArH), 7.59 (br t, J = 7.3 Hz, 1H, ArH), 7.63 (d, J = 8.8 Hz, 2H, ArH), 8.12 (br d, J = 7.3 Hz, 2H, ArH); ¹³C nmr: δ 13.8, 42.4, 55.4, 62.8, 91.8, 114.2, 120.5, 128.61, 128.64, 130.1, 133.4, 133.9, 155.7, 161.5, 169.2, 190.3; HR-MS m/z: Calcd. for C₂₀H₁₉NO₄: 353.1263. Found: 353.1273.

5-Benzoyl-3-(4-chlorophenyl)-4,5-dihydroisoxazole-5-carboxylic Acid Ethyl Ester (**3d-e**).

This compound was obtained as colorless needles (hexane- C_6H_6), mp 108-109°, ir v CO 1755, 1693 cm⁻¹; ¹H nmr: δ 1.16 (t, J = 7.1 Hz, 3H, CH₃), 3.67 (d, J = 17.5 Hz, 1H, CHH), 4.24 (q, J = 7.1 Hz, 2H, OCH₂), 4.54 (d, J = 17.5 Hz, 1H, CHH), 7.37-7.66 (m, 7H, ArH), 8.12 (br d, J = 7.1 Hz, 2H, ArH); ¹³C nmr: δ 13.9, 42.0, 62.9, 92.1, 126.5, 128.6, 129.0, 130.0, 133.1, 134.0, 136.7, 155.1, 168.8, 189.6.

Anal. Calcd. for C₁₉H₁₆ClNO₄: C, 63.78; H, 4.51; N, 3.91. Found: C, 63.70; H, 4.31; N, 3.89. Sep-Oct 2002

5-Benzoyl-3-(4-bromophenyl)-4,5-dihydroisoxazole-5-carboxylic Acid Ethyl Ester (**3d-f**).

This compound was obtained as colorless needles (hexane- C_6H_6), mp 109-110°, ir v CO 1752, 1692 cm⁻¹; ¹H nmr: δ 1.16 (t, J = 7.1 Hz, 3H, CH₃), 3.66 (d, J = 17.6 Hz, 1H, CHH), 4.24 (q, J = 7.1 Hz, 2H, OCH₂), 4.54 (d, J = 17.6 Hz, 1H, CHH), 7.45-7.61 (m, 7H, ArH), 8.12 (br d, J = 7.3 Hz, 2H, ArH); ¹³C nmr: δ 13.9, 42.0, 62.9, 92.1, 125.1, 126.9, 128.4, 128.6, 130.0, 132.0, 133.1, 134.0, 153.2, 168.7, 189.6.

Anal. Calcd. for C₁₉H₁₆BrNO₄: C, 56.73; H, 4.01; N, 3.48. Found: C, 56.96; H, 3.84; N, 3.40.

5-Benzoyl-3-naphthalen-1-yl-4,5-dihydroisoxazole-5-carboxylic Acid Ethyl Ester (**3d-g**).

This compound was obtained as colorless needles (hexane- C_6H_6), mp 97-98°, ir v CO 1750, 1695 cm⁻¹; ¹H nmr: δ 1.18 (t, J = 7.1 Hz, 3H, CH₃), 3.88 (d, J = 17.5 Hz, 1H, CHH), 4.27 (q, J = 7.1 Hz, 2H, OCH₂), 4.79 (d, J = 17.5 Hz, 1H, CHH), 7.46-7.95 (m, 9H, ArH), 8.18 (br d, J = 7.1 Hz, 2H, ArH), 8.96 (br d, J = 8.2 Hz, 1H, ArH); ¹³C nmr: δ 14.0, 45.0, 62.9, 90.8, 124.6, 124.9, 126.4, 126.9, 127.7, 128.3, 128.5, 128.6, 130.0, 130.4, 131.5, 133.3, 133.8, 133.9, 156.7, 169.0, 189.8.

Anal Calcd. for C₂₃H₁₉NO₄: C, 73.98; H, 5.13; N, 3.75. Found: C, 73.92; H, 4.91; N, 3.53.

5-Benzoyl-3-phenyl-4,5-dihydroisoxazole-5-carboxylic Acid Cyclohexyl Ester (**3e-a**).

This compound was obtained as colorless needles (hexane), mp 101°, ir v CO 1754, 1693 cm⁻¹; ¹H nmr: δ 1.20-1.73 (m, 10H, CH₂ x 5), 3.68 (d, J = 17.5 Hz, 1H, CHH), 4.58 (d, J = 17.5 Hz, 1H, CHH), 4.86 (m, 1H, OCH), 7.40-7.73 (m, 8H, ArH), 8.13 (br d, J = 7.3 Hz, 2H, ArH); ¹³C nmr: δ 23.1, 23.3, 25.1, 30.8, 31.1, 42.1, 75.3, 91.9, 126.9, 128.1, 128.5, 128.7, 129.9, 130.6, 133.4, 133.8, 153.9, 168.4, 189.8.

Anal. Calcd. for C₂₃H₂₃NO₄: C, 73.19; H, 6.14; N, 3.71. Found: C, 73.14; H, 6.13; N, 3.60.

5-Benzoyl-3-(4-methylphenyl)-4,5-dihydroisoxazole-5-carboxylic Acid Cyclohexyl Ester (**3e-b**).

This compound was obtained as colorless needles (hexane), mp 101-102°, ir v CO 1751, 1687 cm⁻¹; ¹H nmr: δ 1.21-1.73 (m, 10H, CH₂ x 5), 2.38 (s, 3H, CH₃), 3.64 (d, J = 17.5 Hz, 1H, CHH), 4.55 (d, J = 17.5 Hz, 1H, CHH), 4.86 (m, 1H, OCH), 7.22 (d, J = 8.0 Hz, 2H, ArH), 7.47 (br t, J = 7.3 Hz, 2H, ArH), 7.58 (br t, J = 7.3 Hz, 1H, ArH), 7.59 (d, J = 8.0 Hz, 2H, ArH), 8.13 (br d, J = 7.3 Hz, 2H, ArH); ¹³C nmr: δ 21.6, 23.2, 23.3, 25.1, 30.8, 31.1, 42.3, 75.2, 91.8, 125.2, 126.9, 128.5, 129.4, 129.9, 133.4, 133.7, 141.0, 155.9, 168.4, 189.9.

Anal Calcd. for C₂₄H₂₅NO₄: C, 73.64; H, 6.44; N, 3.58. Found: C, 73.55; H, 6.39; N, 3.60.

5-Benzoyl-3(4-isopropylphenyl)-4,5-dihydroisoxazole-5-carboxylic Acid Cyclohexyl Ester (**3e-c**).

This compound was obtained as colorless needles (hexane), mp 91°, ir v CO 1755, 1690 cm⁻¹; ¹H nmr: δ 1.26 (d, J = 6.9 Hz, 6H, CH₃), 1.21-1.73 (m, 10H, CH₂ x 5), 2.94 (heptet, J = 6.9 Hz, 1H, CH), 3.65 (d, J = 17.5 Hz, 1H, CHH), 4.56 (d, J = 17.5 Hz, 1H, CHH), 4.86 (m, 1H, OCH), 7.27 (d, J = 8.3 Hz, 2H, ArH), 7.47 (br t, J = 7.3 Hz, 2H, ArH), 7.59 (br t, J = 7.3 Hz, 1H, ArH), 7.63 (d, J = 8.3 Hz, 2H, ArH), 8.13 (br d, J = 7.3 Hz, 2H, ArH); ¹³C nmr: δ 23.2, 23.3, 23.8, 25.1, 30.8, 31.1, 34.2, 42.3, 75.2, 77.5, 91.7, 125.6, 126.8, 127.0, 128.5, 129.9, 133.4, 133.7, 151.8, 155.8, 168.4, 189.9.

Anal Calcd. for C₂₆H₂₉NO₄: C, 74.44; H, 6.97; N, 3.34. Found: C, 74.31; H, 6.82; N, 3.28.

5-Benzoyl-3-(4-methoxyphenyl)-4,5-dihydroisoxazole-5-carboxylic Acid Cyclohexyl Ester (**3e-d**).

This compound was obtained as colorless needles (hexane- C_6H_6), mp 106-108°, ir v CO 1750, 1693 cm⁻¹; ¹H nmr: δ 1.21-1.73 (m, 10H, CH₂ x 5), 3.63 (d, J = 17.4 Hz, 1H, CHH), 3.84 (s, 3H, CH₃), 4.54 (d, J = 17.4 Hz, 1H, CHH), 4.86 (m, 1H, OCH), 6.92 (d, J = 8.8 Hz, 2H, ArH), 7.47 (br t, J = 7.3 Hz, 2H, ArH), 7.59 (br t, J = 7.3 Hz, 1H, ArH), 7.64 (d, J = 8.8 Hz, 2H, ArH), 8.13 (br d, J = 7.3 Hz, 2H, ArH); ¹³C nmr: δ 23.2, 23.3, 25.1, 30.8, 31.1, 42.4, 55.4, 75.2, 91.7, 114.1, 120.6, 128.46, 128.54, 129.9, 133.4, 133.7, 155.4, 161.3, 168.5, 190.0; HR-MS m/z: Calcd. for C₂₄H₂₅NO₅: 407.1733. Found: 407.1744.

5-Benzoyl-3-(4-chlorophenyl)-4,5-dihydroisoxazole-5-carboxylic Acid Cyclohexyl Ester (**3e-e**).

This compound was obtained as colorless needles (hexane- C_6H_6), mp 106-108°, ir v CO 1750, 1690 cm⁻¹; ¹H nmr: δ 1.19-1.73 (m, 10H, CH₂ x 5), 3.62 (d, J = 17.5 Hz, 1H, CHH), 4.55 (d, J = 17.5 Hz, 1H, CHH), 4.87 (m, 1H, OCH), 7.39 (d, J = 8.6 Hz, 2H, ArH), 7.47 (t, J = 7.3 Hz, 2H, ArH), 7.60 (br t, J = 7.3 Hz, 1H, ArH), 7.64 (d, J = 8.6 Hz, 2H, ArH), 8.13 (br d, J = 7.3 Hz, 2H, ArH); ¹³C nmr: δ 23.1, 23.3, 25.1, 30.8, 31.1, 41.9, 75.4, 92.1, 126.6, 128.2, 128.5, 129.0, 129.9, 133.3, 133.9, 136.7, 155.0, 168.2, 189.5.

Anal. Calcd. for C₂₃H₂₂ClNO₄: C, 67.07; H, 5.38; N, 3.40. Found: C, 66.87; H, 5.28; N, 3.43.

5-Benzoyl-3-(4-bromophenyl)-4,5-dihydroisoxazole-5-carboxylic Acid Cyclohexyl Ester (**3e-f**).

This compound was obtained as colorless needles (hexane), mp 117-118°, ir v CO 1753, 1692 cm⁻¹; ¹H nmr: δ 1.19-1.73 (m, 10H, CH₂ x 5), 3.52 (d, J = 17.5 Hz, 1H, CHH), 4.55 (d, J = 17.5 Hz, 1H, CHH), 4.87 (m, 1H, OCH), 7.37-7.66 (m, 7H, ArH), 8.12 (br d, J = 7.3 Hz, 2H, ArH); ¹³C nmr: δ 23.1, 23.3, 25.1, 30.8, 31.0, 41.9, 75.4, 92.1, 125.0, 127.0, 128.4, 128.5, 129.9, 132.0, 133.3, 133.9, 155.1, 168.2, 189.5.

Anal. Calcd. for C₂₃H₂₂BrNO₄: C, 60.54; H, 4.86; N, 3.07. Found: C, 60.78; H, 4.67; N, 3.13.

5-Benzoyl-3-naphthalen-1-yl-4,5-dihydroisoxazole-5-carboxylic Acid Cyclohexyl Ester (**3e-g**).

This compound was obtained as colorless needles (hexane- C_6H_6), mp 124-125°, ir v CO 1743, 1690 cm⁻¹; ¹H nmr: δ 1.20-1.75 (m, 10H, CH₂ x 5), 3.83 (d, J = 17.4 Hz, 1H, CHH), 4.81 (d, J = 17.4 Hz, 1H, CHH), 4.90 (m, 1H, OCH), 7.46-7.95 (m, 9H, ArH), 8.19 (br d, J = 7.1 Hz, 2H, ArH), 8.95 (br d, J = 8.4 Hz, 1H, ArH); ¹³C nmr: δ 23.2, 23.3, 25.1, 30.8, 31.1, 44.9, 75.3, 90.7, 124.6, 125.0, 126.4, 126.9, 127.6, 128.2, 128.47, 128.54, 129.9, 130.4, 131.1, 133.5, 133.8, 156.6, 168.5, 189.8.

Anal. Calcd. for $C_{27}H_{25}NO_4$: C, 75.86; H, 5.89; N, 3.28. Found: C, 75.63; H, 5.69; N, 3.29.

5-Benzoyl-3-phenyl-4,5-dihydroisoxazole-5-carboxylic Acid Benzyl Ester (**3f-a**).

This compound was obtained as colorless needles (hexane-C₆H₆), mp 103-104°, ir v CO 1748, 1685 cm⁻¹; ¹H nmr: δ 3.66

(d, J = 17.6 Hz, 1H, CHH), 4.57 (d, J = 17.6 Hz, 1H, CHH), 5.19 (ABq, J = 12.1 Hz, 2H, OCH₂Ph), 7.12-7.71 (m, 13H, ArH), 8.05 (br d, J = 7.1 Hz, 2H, ArH); 13 C nmr: δ 42.1, 68.2, 91.9, 127.1, 128.0, 128.3, 128.53, 128.57, 128.62, 128.8, 130.0, 130.8, 133.2, 134.0, 134.3, 156.1, 168.9, 189.6.

Anal Calcd. for C₂₄H₁₉NO₄: C, 74.79; H, 4.97; N, 3.63. Found: C, 74.68; H, 4.89; N, 3.33.

5-Benzoyl-3-(4-methylphenyl)-4,5-dihydroisoxazole-5-carboxylic Acid Benzyl Ester (**3f-b**).

This compound was obtained as colorless needles (hexane- C_6H_6), mp 115°, ir v CO 1748, 1686 cm⁻¹; ¹H nmr: δ 2.38 (s, 3H, CH₃), 3.65 (d, J = 17.6 Hz, 1H, CHH), 4.54 (d, J = 17.6 Hz, 1H, CHH), 5.19 (ABq, J = 12.1 Hz, 2H, OCH₂Ph), 7.10-7.60 (m, 12H, ArH), 8.05 (br d, J = 7.2 Hz, 2H, ArH); ¹³C nmr: δ 21.5, 42.3, 68.1, 91.7, 125.2, 127.0, 128.3, 128.53, 128.54, 128.6, 129.5, 130.0, 133.9, 134.3, 141.2, 156.1, 169.0, 189.7.

Anal. Calcd. for C₂₅H₂₁NO₄: C, 75.17; H, 5.30; N, 3.51. Found: C, 74.99; H, 5.18; N, 3.41.

5-Benzoyl-3-(4-isopropylphenyl)-4,5-dihydroisoxazole-5-carboxylic Acid Ethyl Ester (**3f-c**).

This compound was obtained as colorless needles (hexane), mp 116-117°, ir v CO 1757, 1696 cm⁻¹; ¹H nmr: δ 1.25 (d, J = 6.9 Hz, 6H, CH₃), 2.93 (heptet, J = 6.9 Hz, 1H, CH), 3.65 (d, J = 17.6 Hz, 1H, CHH), 4.54 (d, J = 17.6 Hz, 1H, CHH), 5.19 (ABq, J = 12.1 Hz, 2H, OCH₂Ph), 7.12-7.63 (m, 12H, ArH), 8.05 (br d, J = 7.3 Hz, 2H, ArH); ¹³C nmr: δ 23.7, 34.1, 42.3, 68.1, 91.7, 125.5, 126.9, 127.1, 128.3, 128.53, 128.54, 128.6, 130.0, 133.3, 133.9, 134.3, 152.0, 156.1, 169.0, 189.8.

Anal Calcd. for C₂₇H₂₅NO₄: C, 75.86; H, 5.89; N, 3.28. Found: C, 75.79; H, 5.87; N, 3.24.

5-Benzoyl-3-(4-methoxyphenyl)-4,5-dihydroisoxazole-5-carboxylic Acid Benzyl Ester (**3f-d**).

This compound was obtained as colorless needles (hexane- C_6H_6), mp 119-120°, ir v CO 1749, 1690 cm⁻¹; ¹H nmr: δ 3.64 (d, J = 17.6 Hz, 1H, CHH), 3.84 (s, 3H, CH₃), 4.53 (d, J = 17.6 Hz, 1H, CHH), 5.19 (ABq, J = 12.3 Hz, 2H, OCH₂Ph), 6.92 (d, J = 8.9 Hz, 2H, ArH), 7.13-7.26 (m, 5H, ArH), 7.39 (br t, J = 7.3 Hz, 2H, ArH), 7.52 (br t, J = 7.3 Hz, 1H, ArH), 7.62 (d, J = 8.9 Hz, 2H, ArH), 8.05 (br d, J = 7.3 Hz, 2H, ArH); ¹³C nmr: δ 42.4, 55.4, 68.1, 77.2, 91.7, 114.2, 120.5, 128.3, 128.5, 128.6, 128.7, 130.1, 133.3, 133.9, 134.4, 155.7, 161.6, 169.1, 189.8; HR-MS m/z: Calcd. for $C_{20}H_{19}NO_4$: 415.1420. Found: 415.1425.

5-Benzoyl-3-(4-chlorophenyl)-4,5-dihydroisoxazole-5-carboxylic Acid Benzyl Ester (**3f-e**).

This compound was obtained as colorless needles (hexane- C_6H_6), mp 117-118°, ir v CO 1747, 1685 cm⁻¹; ¹H nmr: δ 3.62 (d, J = 17.5 Hz, 1H, CHH), 4.54 (d, J = 17.5 Hz, 1H, CHH), 5.19 (ABq, J = 12.0 Hz, 2H, OCH₂Ph), 7.15-7.63 (m, 8H, ArH), 7.38, (d, J = 8.7 Hz, 2H, ArH), 7.61, (d, J = 8.7 Hz, 2H, ArH), 8.04 (br d, J = 7.1 Hz, 2H, ArH); ¹³C nmr: δ 42.0, 68.3, 92.0, 126.4, 128.2, 128.46, 128.52, 128.6, 129.0, 129.9, 133.0, 133.9, 134.1, 136.7, 155.1, 168.6, 189.1.

Anal. Calcd. for C₂₄H₁₈ClNO₄: C, 68.66; H, 4.32; N, 3.34. Found: C, 68.66; H, 4.21; N, 3.23.

5-Benzoyl-3-(4-bromophenyl)-4,5-dihydroisoxazole-5-carboxylic Acid Benzyl Ester (**3f-f**).

This compound was obtained as colorless needles (hexane- C_6H_6), mp 134-135°, ir v CO 1743, 1685 cm⁻¹; ¹H nmr δ : 3.61 (d, J = 17.6 Hz, 1H, CHH), 4.54 (d, J = 17.6 Hz, 1H, CHH), 5.19 (ABq, J = 12.1 Hz, 2H, OCH₂Ph), 7.14-7.55 (m, 12H, ArH), 8.04 (br d, J = 7.4 Hz, 2H, ArH); ¹³C nmr: δ 41.9, 68.3, 92.0, 125.1, 126.9, 128.2, 128.4, 128.46, 128.52, 128.6, 129.9, 132.0, 133.0, 133.9, 134.1, 155.2, 168.6, 189.1.

Anal. Calcd. for C₂₄H₁₈BrNO₄: C, 62.08; H, 3.91; N, 3.02. Found: C, 62.36; H, 3.76; N, 2.91.

5-Benzoyl-3-naphthalen-1-yl-4,5-dihydroisoxazole-5-carboxylic Acid Benzyl Ester (**3f-g**).

This compound was obtained as colorless needles (hexane- C_6H_6), mp 99-101°, ir v CO 1754, 1685 cm⁻¹; ¹H nmr: δ 3.84 (d, J = 17.5 Hz, 1H, CHH), 4.79 (d, J = 17.5 Hz, 1H, CHH), 5.22 (s, 2H, OCH₂Ph), 7.17-7.94 (m, 14H, ArH), 8.11 (br d, J = 7.3 Hz, 2H, ArH), 8.92 (br d, J = 8.2 Hz, 1H, ArH); ¹³C nmr: δ 45.0, 68.2, 90.7, 124.6, 124.8, 126.4, 126.8, 127.7, 128.2, 128.3, 128.46, 128.48, 128.6, 130.0, 130.4, 131.5, 133.2, 133.8, 133.9, 134.2, 156.3, 168.9, 189.4.

Anal Calcd. for C₂₈H₂₁NO₄: C, 77.23; H, 4.86; N, 3.22. Found: C, 77.15; H, 4.77; N, 3.14.

REFERENCES AND NOTES

[1] Reviews, see; A. Padwa, In Comprehensive Organic Synthesis, ed. by B. M. Trost, I. Fleming, Pergamon Press, Oxford, 1991, Vol. **4**, pp 1069-1109; W. Carruthers, Cycloaddition Reactions in Organic Synthesis, Pergamon Press, Tokyo, 1990, pp 269-331; C. J. Easton C. M. M. Hughes, G. P. Savage and G. W. Simpson, In Advances in Heterocyclic Chemistry, ed. by A. R. Katritzky, Academic Press, New York, 1994, Vol. **60**, pp 261-328.

[2] D. P. Curran, In Advance in Cycloaddition, ed. by D. P. Curran, JAI Press, London, 1988, Vol 1, pp 129-189.

[3] V. Jäger and V. Müller, *Tetrahedron Letters*, **23**, 4777 (1982); V. Jäger, V. Müller and E. F. Paulus, *Tetrahedron Letters*, **26**, 2997 (1985).

[4a] K. N. Houk, J. Sims, R. E. Duke, Jr., R. W. Strozier and J. K. George, J. Am. Chem. Soc., 95, 7287 (1973); [b] K. N. Houk, J. Sims, C. R. Watts and L. J. Luskus, J. Am. Chem. Soc., 95, 7301 (1973).

[5a] M. Yamauchi, S. Katayama and T. Watanabe, Synthesis,
935 (1982); [b] M. Yamauchi, S. Katayama, O. Baba and T. Watanabe, J. Chem. Soc., Perkin Trans. 1, 3041 (1990); [c] M. Yamauchi, Y. Honda, N. Matsuki, T. Watanabe, T. Date and H. Hiramatsu, J. Org. Chem., 61, 2719 (1996); [d] M. Yamauchi and M. Yajima, Pharm. Bull., 49, 1638 (2001); [e] M. Yamauchi, T. Aoki,
M.-Z. Li and Y. Honda, Tetrahedron: Asymmetry, 12, 3113 (2001).

[6] K. E. Larsen and K. B. G. Torssell, *Tetrahedron*, **40**, 2985 (1984).

[7] M. Yamauchi, unpublished results.

[8] M. Christl and R. Huisgen, Chem. Ber., 106, 3345 (1973).