

## 3-Acyl-1,2,3,4-tetrahydropyridine-2,4-diones: Synthesis and Chemical Properties

D. B. Rubinov, T. A. Zheldakova, I. L. Rubinova, and A. V. Baranovskii

*Institute of Bioorganic Chemistry, National Academy of Sciences of Belarus,  
ul. Akademika Kuprevicha 5/2, Minsk, 220141 Belarus  
e-mail: rub\_irin@mail.ru*

Received May 10, 2006

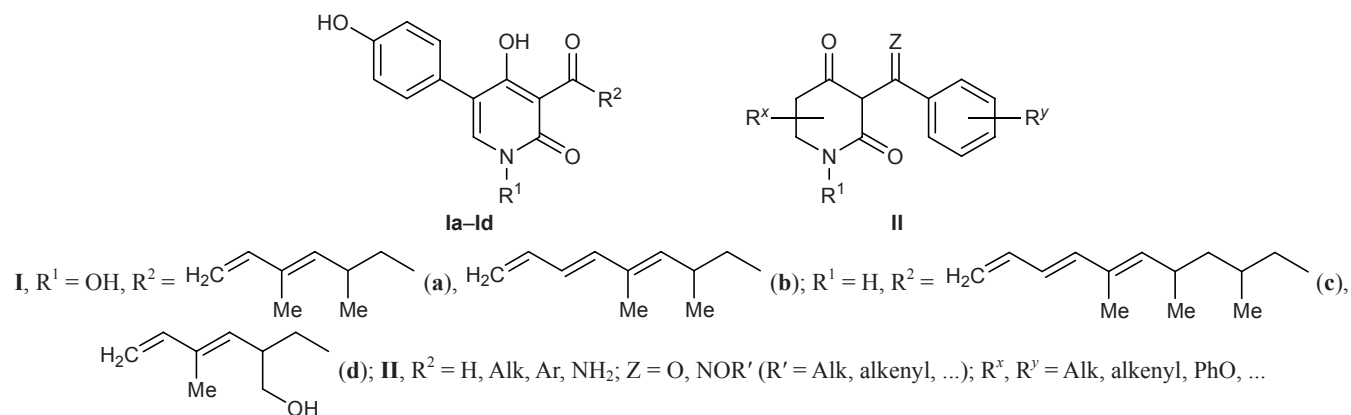
**Abstract**—N-Substituted 6-methyl-1,2,3,4-tetrahydropyridine-2,4-diones reacted with aliphatic carboxylic acid chlorides in the presence of pyridine or triethylamine to give the corresponding 4-O-acyl derivatives which underwent O,C-migration of the acyl group by the action of 2 equiv of triethylamine and a catalytic amount of 2-hydroxy-2-methylpropanenitrile. Reactions of 3-acyl-6-methyl-1,2,3,4-tetrahydropyridine-2,4-diones thus formed with aliphatic and aromatic amines gave the corresponding enamino derivatives at the side acyl group. Enamino derivatives at the C<sup>4</sup>=O group were obtained by transformation of 3-acyl-1,2,3,4-tetrahydropyridine-2,4-diones into 3-acyl-4-methoxy-6-methyl-1,2-dihydropyridin-2-ones via alkylation with dimethyl sulfate and subsequent treatment with amines.

**DOI:** 10.1134/S1070428008030196

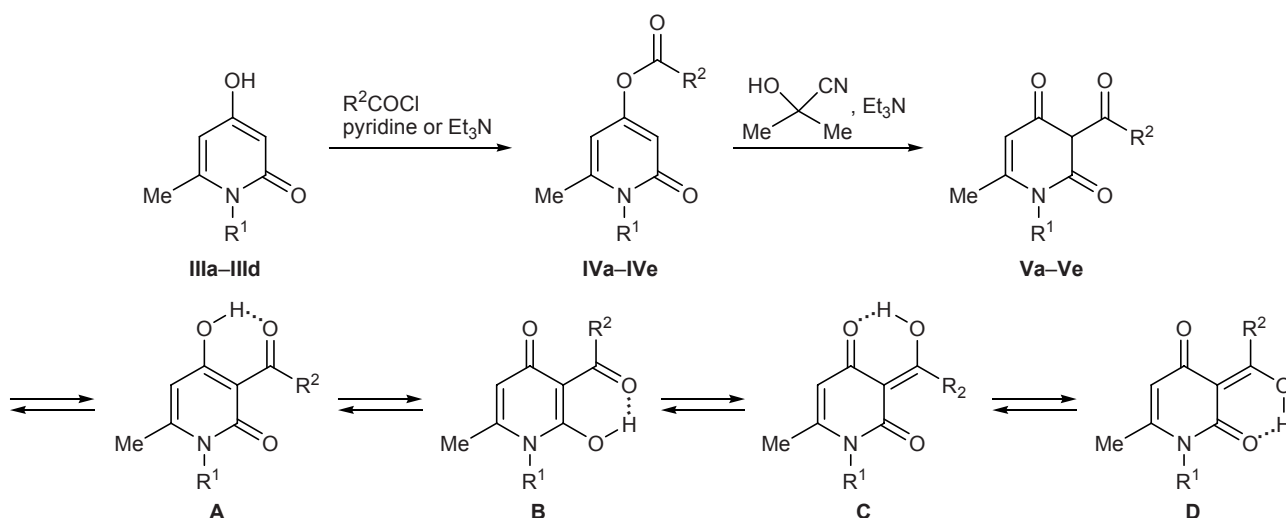
Pyridinone and pyridinedione derivatives are widely used in the synthesis of natural [1] and biologically active compounds [2]. 3-Acyl-2,4-dioxypyridine fragment constitutes a structural base of Tenellin (**Ia**), Bassianin (**Ib**) [3], Militarone (**Ic**) [4], Pyridovericin (**Id**) [5], and other natural alkaloid pigments isolated from entomopathogenic fungi. It is known that synthetic 3-aroil-substituted pyridine-2,4-diones and oximes derived therefrom (**II**, Z = NOR') exhibit herbicidal activity [6–9]; like carbocyclic β-triketones, they are prepared from the corresponding pyridinediones via reaction with aroyl chlorides, followed by O–C migration of the aroyl group in enol esters in the presence of potassium cyanide [10, 11]. Syntheses of β-triketones

of the pyrimidine series having an aliphatic side chain include a number of steps, and the yields of the target products are often poor [4, 12–14]. Acylation of pyridine-2,4-diones with aliphatic carboxylic acid chlorides was not reported.

We previously proposed a simple and efficient procedure for the synthesis of 1,6-disubstituted pyridine-2,4-diones **III** from 5-(1,3-dioxobutyl)-2,2-dimethyl-1,3-dioxane-4,6-dione [15]. While continuing studies on the chemical properties of pyridine-2,4-dione derivatives, we have found that the procedure used by us previously for the preparation of 2-aroilcyclohexane-1,3-diones [16] ensures synthesis of 3-acylpyridine-2,4-diones **V** in good yields. Our results contradict the



Scheme 1.



III, R<sup>1</sup> = Me (a), Pr (b), PhCH<sub>2</sub> (c), 4-MeOC<sub>6</sub>H<sub>4</sub> (d); IV, V, R<sup>1</sup> = R<sup>2</sup> = Me (a); R<sup>1</sup> = Pr, R<sup>2</sup> = Et (b); R<sup>1</sup> = PhCH<sub>2</sub>, R<sup>2</sup> = Et (c), Me (f); R<sup>1</sup> = 4-MeOC<sub>6</sub>H<sub>4</sub>, R<sup>2</sup> = Pr (d); R<sup>1</sup> = Me, R<sup>2</sup> = Bu (e).

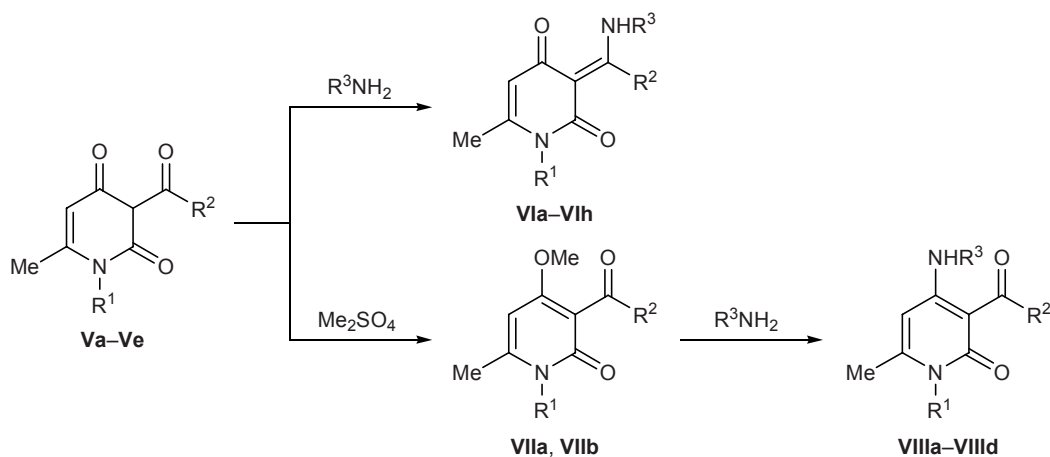
data of Schnell and Kappe [11] who failed to obtain 3-acylpyridine-2,4-diones following an analogous approach. It should be noted that we did not succeed in synthesizing  $\beta$ -triketones **V** using Lewis acids and 4-dimethylaminopyridine as catalysts.

Treatment of diketones **IIIa–IIIId** with acetyl, propionyl, and butyryl chlorides in methylene chloride in the presence of pyridine gave enol esters **IVa–IVf** with high regioselectivity. Compounds **IVa–IVf** can be isolated as individual substances in 90–95% yield or subjected (without isolation) to O–C-migration of the acyl group with formation of 3-acyl-1,2,3,4-tetrahydropyri-

dine-2,4-diones **Va–Vf** in the presence of excess triethylamine and 2-hydroxy-2-methylpropanenitrile as catalyst (Scheme 1).

The assumed structure of enol esters **IV** follows from the <sup>1</sup>H NMR spectra of compounds **IVa–IVc** and **IVf**, where the 3-H and 5-H olefinic protons ( $\delta$  5.50–6.50 ppm) displayed an allylic coupling constant <sup>4</sup>J of 2.0–2.5 Hz. Like other cyclic  $\beta$ -triketones [17], 3-acylpyridine-2,4-diones **Va–Vf** are completely enolized: their <sup>1</sup>H NMR spectra contain a one-proton signal from the chelated hydroxy group in the region  $\delta$  15–16 ppm. Theoretically, unsymmetrical heterocyclic  $\beta$ -triketones

Scheme 2.



VI, R<sup>1</sup> = R<sup>2</sup> = Me, R<sup>3</sup> = Ph (a), CH<sub>2</sub>=CHCH<sub>2</sub> (b); R<sup>1</sup> = Me, R<sup>2</sup> = Bu, R<sup>3</sup> = CH<sub>2</sub>=CHCH<sub>2</sub> (c), 4-MeOC<sub>6</sub>H<sub>4</sub> (d, h); R<sup>1</sup> = Pr, R<sup>2</sup> = Et, R<sup>3</sup> = PhCH<sub>2</sub> (e); R<sup>1</sup> = 4-MeOC<sub>6</sub>H<sub>4</sub>, R<sup>2</sup> = Pr, R<sup>3</sup> = PhCH<sub>2</sub> (f), 4-MeOC<sub>6</sub>H<sub>4</sub> (g); R<sup>1</sup> = Me, R<sup>2</sup> = Bu, R<sup>3</sup> = 4-MeOC<sub>6</sub>H<sub>4</sub> (h); VII, R<sup>1</sup> = Me, R<sup>2</sup> = Bu (a); R<sup>1</sup> = 4-MeOC<sub>6</sub>H<sub>4</sub>, R<sup>2</sup> = Pr (b); VIII, R<sup>1</sup> = 4-MeOC<sub>6</sub>H<sub>4</sub>, R<sup>2</sup> = Pr, R<sup>3</sup> = PhCH<sub>2</sub> (a), 4-MeOC<sub>6</sub>H<sub>4</sub> (b); R<sup>1</sup> = Me, R<sup>2</sup> = Bu, R<sup>3</sup> = 4-MeOC<sub>6</sub>H<sub>4</sub> (c), CH<sub>2</sub>=CHCH<sub>2</sub> (d).

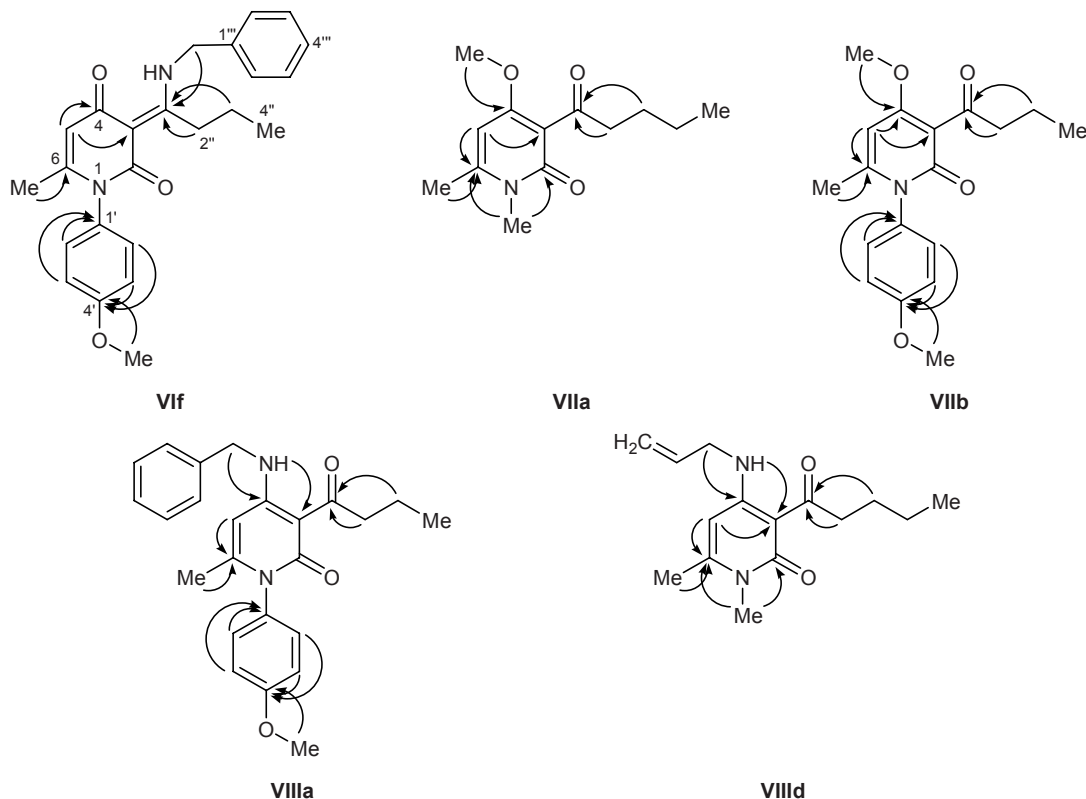
**Va–Vf** can give rise to four tautomeric forms **A–D**. The presence of only one enol proton signal suggests that compounds **Va–Vf** have structure **A** or **C**; the contribution of tautomers **B** and **D** is quite insignificant, or they are absent at all.

3-Acylpyridinediones **Va–Vf**, as well as other cyclic  $\beta$ -triketones, reacted with amines at the acyl carbonyl group, yielding enamines **Via–Vih** (Scheme 2). The reactions with aliphatic amines (allylamine and benzylamine) in boiling toluene required several hours, whereas the conversion of triketones **V** in the reactions with aromatic amines (aniline and *p*-methoxyaniline) was not complete even after heating for a weak, and the yields of the corresponding enamines **VI** did not exceed 50%.

With a view to extend the synthetic potential of 3-acylpyridine-2,4-diones we tried to obtain their enamino derivatives at one of the endocyclic carbonyl groups. Analogous transformations of carbocyclic  $\beta$ -triketones are usually accomplished via intermediate preparation of the corresponding enol methyl ethers, and these reactions involve no difficulties [17]. We previously synthesized regioisomeric enol methyl ethers at both endocyclic carbonyl groups of sulfur-containing heterocyclic  $\beta$ -triketones of the 3-acylthiotetronic acid

series [18] and 3-acetyltetrahydrothiopyran-2,4-dione [19]. 4-Acyl-2*H*-thiopyran-3,5-diones reacted with methylating agents to give complex mixtures of products, from which we failed to isolate the desired enol ethers [20]. By heating triketones **Vc** and **Vf** with 1.2 equiv of dimethyl sulfate and 4 equiv of anhydrous potassium carbonate in toluene we obtained the corresponding 4-methoxy derivatives **VIIa** and **VIIb** in almost quantitative yield. Compounds **VIIa** and **VIIb** reacted with allylamine, benzylamine, aniline, and *p*-methoxyaniline to produce enamino derivatives **VIIIa–VIIId** at the  $C^4=O$  carbonyl group. Unlike enol ethers derived from  $\beta$ -triketones of the cyclohexane [17] and heterocyclic series [18, 19], 4-methoxypyridinones **VIIa** and **VIIb** turned out to be low reactive; as in the synthesis of enamino derivatives **Via–Vih** at the side-chain carbonyl group, the yields in the reactions of **VIIa** and **VIIb** with aromatic amines were considerably lower.

The product structure was determined using two-dimensional NMR techniques (COSY, HSQC, HMBC), and complete assignment of carbon signals in the  $^{13}C$  NMR spectra was made. The assignment of signals from carbon atoms attached to protons involved no difficulties, for the COSY and HSQC data were fully



Principal  $^{13}C$ - $^1H$  interactions in the HMBC spectra of compounds **VI f**, **VII a**, **VII b**, **VIII a**, and **VIII d**.

consistent with each other. Some problems appeared while assigning signals from protons in the 4'-methoxyphenyl substituent in compounds **Vd**, **Vif**, **VIIb**, and **VIIIa**. The COSY spectra of these compounds displayed only a weak cross-peak between the methyl protons and 3'-H. Nevertheless, taking into account more upfield positions of the  $^{13}\text{C}$  ( $\delta_{\text{C}}$  114–115 ppm) and  $^1\text{H}$  signals ( $\delta$  6.98–7.03 ppm), they were assigned to  $\text{C}^{3'}$  and 3'-H, respectively, while the signals located at  $\delta_{\text{C}}$  129–130 ppm and  $\delta$  7.07–7.09 ppm were assigned to  $\text{C}^{2'}$  and 2'-H.

Signals from quaternary carbon atoms were assigned by analysis of long-range  $^1\text{H}$ – $^{13}\text{C}$  couplings in the HMBC spectra. The principal interactions are shown in figure. All compounds having an exocyclic carbonyl group displayed cross-peaks between the carbonyl carbon atoms ( $\delta_{\text{C}}$  203–208 ppm) and protons in the  $\alpha$ - and  $\beta$ -positions with respect to the carbonyl group. The corresponding protons in molecule **Vif** interact with the carbon nucleus resonating at  $\delta_{\text{C}}$  179.37 ppm, and that carbon nucleus also gives a cross-peak with methylene protons in the benzyl group. These data unambiguously indicate that the enamino fragment originates from the exocyclic carbonyl group. The lactam carbonyl carbon atom ( $\delta_{\text{C}}$  162–165 ppm) showed no strong cross-peaks because of the absence of protons in the vicinity of  $\text{C}^2=\text{O}$ . Only the HMBC spectra of *N*-methyl derivatives **VIIa** and **VIIIc** contained cross peaks resulting from coupling between the methyl protons and the carbonyl carbon atom. The quaternary carbon atom in the 3-position of the pyridine ring is readily identified taking into account upfield position of the corresponding signal ( $\delta_{\text{C}}$  101–112 ppm) and the presence of a cross-peak due to coupling with 5-H. Enamines **VIIIa** and **VIIIc** also displayed a cross-peak between  $\text{C}^3$  and NH proton. It should be noted that the  $\text{C}^3$  signal in the spectra of methyl ethers **VIIa** and **VIIb** is observed in a weaker field as compared to the other examined compounds ( $\Delta\delta_{\text{C}} \approx 10$  ppm).

The  $\text{C}^4$  signal was identified by the presence of a weak cross peak with 5-H and (for 4-substituted compounds) by the coupling with protons of the methoxy group (**VIIa**, **VIIb**) and methylene protons in the secondary amine moiety (**VIIIa**, **VIIIc**). Here, variations of  $^{13}\text{C}$  chemical shifts reflect the character of substitution and degree of conjugation in the system. The  $\text{C}^4$  signal in the spectra of enamino derivatives appears at  $\delta_{\text{C}}$  159 ppm, methoxy compounds display the  $\text{C}^4$  signal in a weaker field (by 5–6 ppm), while the  $\text{C}^4$  nucleus in diketone **Vif** resonates at  $\delta_{\text{C}}$  183.73 ppm.

Interactions with protons in the methyl group and 5-H allowed us to distinguish signal from the quaternary  $\text{C}^6$  atom in the pyridine ring. *N*-Methyl derivatives **VIIa** and **VIIIc** additionally showed a cross-peak between  $\text{C}^6$  and  $\text{NCH}_3$  protons. Signals from the quaternary carbon atom in the methoxyphenyl substituent in compounds **Vif**, **VIIb**, and **VIIIa** were assigned on the basis of coupling between the  $\text{OCH}_3$  protons and carbon nucleus resonating at  $\delta_{\text{C}}$  159 ppm; the chemical shift of the latter was also taken into account.

Unlike carbo- [21] and heterocyclic  $\beta$ -triketones [22, 23] studied previously, we failed to obtain condensation products of triketones **IIIa** and **IIIb** with isoquinoline derivatives (diaz analogs of steroids) regardless of the reaction conditions (acid or base catalysis).

## EXPERIMENTAL

The IR spectra of solid products were recorded in KBr on a UR-20 spectrometer; liquid products were examined as films (neat). The  $^1\text{H}$  and  $^{13}\text{C}$  NMR spectra were measured on a Bruker Avance 500 instrument at 500 and 125 MHz, respectively; chloroform-*d* was used as solvent, and tetramethylsilane, as reference. The mass spectra were run on an MKh-1320 spectrometer. The melting points were determined on a Boetius hot stage. The progress of reactions and the purity of products were monitored by thin-layer chromatography on Silufol UV-254 or Alufol UV-254 plates (Merck); spots were detected under UV light, followed by spraying with a solution of iron(III) chloride. Preparative thin-layer chromatography was performed using silica gel Kieselgel 60 HF<sub>254</sub> (Merck).

**Enol esters IVa–IVd (general procedure).** Pyridine, 10.5 ml (0.13 mol), and the corresponding carboxylic acid chloride, 0.11 mol, were added under stirring to a mixture of 0.1 mol of diketone **IIIa–IIIc** and 100 ml of methylene chloride. The mixture was stirred for 24 h at room temperature (TLC), 50 ml of cold water was added, the mixture was acidified to pH 5 by adding 2 N hydrochloric acid, the organic phase was separated, washed with water and a 5% solution of sodium carbonate, and dried over magnesium sulfate, and the solvent was removed on a rotary evaporator.

**1,6-Dimethyl-2-oxo-1,2-dihydropyridin-4-yl acetate (IVa).** Yield 95%, mp 136–137°C (from diethyl ether). IR spectrum,  $\nu$ ,  $\text{cm}^{-1}$ : 1775, 1670, 1600, 1570.  $^1\text{H}$  NMR spectrum,  $\delta$ , ppm: 2.27 s (3H, 6- $\text{CH}_3$ ), 2.36 s (3H,  $\text{CH}_3\text{CO}$ ), 3.50 s (3H,  $\text{NCH}_3$ ), 5.96 d (1H, 5-H,  $J = 2.5$  Hz), 6.23 d (1H, 3-H,  $J = 2.5$  Hz). Found,

%, C 59.80; H 6.21; N 7.87.  $[M]^+$  181.  $C_9H_{11}NO_3$ . Calculated, %: C 59.66; H 6.12; N 7.73.  $M$  181.19.

**6-Methyl-2-oxo-1-propyl-1,2-dihydropyridin-4-yl propanoate (IVb).** Yield 90%. Oily substance. IR spectrum,  $\nu$ ,  $cm^{-1}$ : 1780, 1670, 1600, 1570.  $^1H$  NMR spectrum,  $\delta$ , ppm: 0.98 t (3H,  $CH_3CH_2CH_2$ ,  $J = 7.0$  Hz), 1.23 t (3H,  $CH_3CH_2CO$ ,  $J = 7.5$  Hz), 1.72 m (2H,  $CH_3CH_2CH_2$ ), 2.38 s (3H, 6- $CH_3$ ), 2.54 q (2H,  $CH_3CH_2CO$ ,  $J = 7.5$  Hz), 3.96 m (2H,  $NCH_2$ ), 5.94 d (1H, 5-H,  $J = 2.5$  Hz), 6.20 d (1H, 3-H,  $J = 2.5$  Hz). Found, %: C 64.60; H 7.83; N 6.43.  $[M]^+$  223.  $C_{12}H_{17}NO_3$ . Calculated, %: C 64.55; H 7.67; N 6.27.  $M$  223.27.

**1-Benzyl-6-methyl-2-oxo-1,2-dihydropyridin-4-yl propanoate (IVc).** Yield 93%, mp 86–87°C. IR spectrum,  $\nu$ ,  $cm^{-1}$ : 1765, 1660, 1600, 1580.  $^1H$  NMR spectrum,  $\delta$ , ppm: 1.25 t (3H,  $CH_3CH_2$ ,  $J = 7.5$  Hz), 2.28 s (1H, 6- $CH_3$ ), 2.58 q (2H,  $CH_3CH_2$ ,  $J = 7.5$  Hz), 5.34 s (2H,  $CH_2C_6H_5$ ), 5.95 d (1H, 5-H,  $J = 2.5$  Hz), 6.33 d (1H, 3-H,  $J = 2.5$  Hz), 7.13–7.37 m (5H,  $C_6H_5$ ). Found, %: C 70.96; H 6.51; N 5.20.  $[M]^+$  271.  $C_{16}H_{17}NO_3$ . Calculated, %: C 70.83; H 6.32; N 5.16.  $M$  271.32.

**1-(4-Methoxyphenyl)-6-methyl-2-oxo-1,2-dihydropyridin-4-yl butanoate (IVd).** Yield 97%, mp 109–110°C. IR spectrum,  $\nu$ ,  $cm^{-1}$ : 1780, 1670, 1610, 1570.  $^1H$  NMR spectrum,  $\delta$ , ppm: 1.04 t (3H,  $CH_3CH_2$ ,  $J = 7.7$  Hz), 1.77 m (2H,  $CH_3CH_2$ ), 1.97 s (3H, 6- $CH_3$ ), 2.53 t (2H,  $CH_2CH_2$ ,  $J = 7.3$  Hz), 3.85 s (3H,  $OCH_3$ ), 6.00 d (1H, 5-H,  $J = 2.2$  Hz), 6.28 d (1H, 3-H,  $J = 2.2$  Hz), 7.03 d (2H,  $C_6H_4$ ,  $J = 8.8$  Hz), 7.10 d (2H,  $C_6H_4$ ,  $J = 8.8$  Hz). Found, %: C 67.69; H 6.47; N 4.48.  $[M]^+$  301.  $C_{17}H_{19}NO_4$ . Calculated, %: C 67.76; H 6.36; N 4.65.  $M$  301.34.

**3-Acyl-6-methyl-1,2,3,4-tetrahydropyridine-2,4-diones Va–Vd (general procedure).** Compound **IVa–IVd**, 0.05 mol, was dissolved in 50 ml of methylene chloride, 14 ml (0.1 mol) of triethylamine and 0.5 ml (0.005 mol) of 2-hydroxy-2-methylpropanenitrile were added, and the mixture was stirred for 48 h at 25–30°C, following the disappearance of initial compound **IVa–IVd** (TLC). The mixture was acidified to pH 5 with 10% hydrochloric acid, and the organic phase was separated, washed with water, dried over magnesium sulfate, and passed through a thin layer of silica gel. The solvent was removed under reduced pressure on a rotary evaporator, and the oily residue was recrystallized from ethyl acetate–petroleum ether.

**3-Acetyl-1,6-dimethyl-1,2,3,4-tetrahydropyridine-2,4-dione (Va).** Yield 83%, mp 130–131°C. IR

spectrum,  $cm^{-1}$ : 1655 (C=O, lactam), 1615 ( $CH_3C=O$ ), 1580 (C=C, enol).  $^1H$  NMR spectrum,  $\delta$ , ppm: 2.36 s (3H, 6- $CH_3$ ), 2.74 s (3H,  $COCH_3$ ), 3.45 s (3H,  $CH_3N$ ), 5.86 s (1H, 5-H), 15.52 s (1H, OH). Found, %: C 59.59; H 6.07; N 7.84.  $[M]^+$  181.  $C_9H_{11}NO_3$ . Calculated, %: C 59.66; H 6.12; N 7.73.  $M$  181.19.

**6-Methyl-1-propyl-3-propionyl-1,2,3,4-pyridine-2,4-dione (Vb).** Yield 85%, mp 58–59°C. IR spectrum,  $\nu$ ,  $cm^{-1}$ : 1665 (C=O, lactam), 1620 (EtC=O), 1570 (C=C, enol).  $^1H$  NMR spectrum,  $\delta$ , ppm: 0.99 t (3H,  $CH_3CH_2CH_2$ ,  $J = 7.4$  Hz), 1.16 t (3H,  $CH_3CH_2CO$ ,  $J = 7.4$  Hz), 1.69 m (2H,  $CH_3CH_2CH_2$ ), 2.37 s (3H, 6- $CH_3$ ), 3.18 q (2H,  $CH_3CH_2CO$ ,  $J = 7.0$  Hz), 3.87 m (2H,  $NCH_2$ ), 5.81 s (1H, 5-H), 15.60 s (1H, OH). Found, %: C 64.47; H 7.50; N 6.37.  $[M]^+$  223.  $C_{12}H_{17}NO_3$ . Calculated, %: C 64.55; H 7.67; N 6.27.  $M$  223.27.

**1-Benzyl-6-methyl-3-propionyl-1,2,3,4-tetrahydropyridine-2,4-dione (Vc).** Yield 80%, mp 75–76°C. IR spectrum,  $\nu$ ,  $cm^{-1}$ : 1670 (C=O, lactam), 1625 (EtC=O), 1570 (C=C, enol).  $^1H$  NMR spectrum,  $\delta$ , ppm: 1.60 t (3H,  $CH_3CH_2$ ,  $J = 7.0$  Hz), 2.28 s (3H, 6- $CH_3$ ), 3.21 q (2H,  $CH_3CH_2$ ), 5.27 s (2H,  $CH_2C_6H_5$ ), 5.86 s (1H, 5-H), 7.13 d (2H,  $H_{arom}$ ), 7.26 m (1H,  $H_{arom}$ ), 7.33 m (2H,  $H_{arom}$ ), 15.74 s (1H, OH). Found, %: C 70.89; H 6.33; N 5.22.  $[M]^+$  271.  $C_{16}H_{17}NO_3$ . Calculated, %: C 70.83; H 6.32; N 5.16.  $M$  271.32.

**3-Butyryl-1-(4-methoxyphenyl)-6-methyl-1,2,3,4-tetrahydropyridine-2,4-dione (Vd).** Yield 82%, mp 105–106°C. IR spectrum,  $\nu$ ,  $cm^{-1}$ : 1670 (C=O, lactam); 1620 (PrC=O); 1610, 1570 (C=C, enol).  $^1H$  NMR spectrum,  $\delta$ , ppm: 0.94 t (3H,  $CH_3CH_2$ ,  $J = 7.4$  Hz), 1.66 m (2H,  $CH_3CH_2$ ), 1.99 s (3H,  $CH_3CH_2$ ), 2.28 s (3H,  $CH_3C=CH$ ), 3.11 t (2H,  $CH_2CH_2$ ,  $J = 7.4$  Hz), 3.85 s (3H,  $OCH_3$ ), 5.93 s (1H, 5-H), 7.03 m (2H,  $H_{arom}$ ), 7.08 m (2H,  $H_{arom}$ ), 15.97 s (1H, OH).  $^{13}C$  NMR spectrum,  $\delta_C$ , ppm: 13.82 ( $C^{4''}$ ), 17.36 ( $C^{3''}$ ), 22.33 (6-Me), 44.71 ( $C^{2''}$ ), 55.54 (MeO), 100.90 ( $C^5$ ), 105.43 ( $C^3$ ), 115.12 ( $C^{3'}$ ), 129.08 ( $C^{2'}$ ), 130.60 ( $C^{1'}$ ), 154.08 ( $C^6$ ), 159.72 ( $C^{4'}$ ), 163.50 ( $C^2$ ), 176.30 ( $C^4$ ), 208.30 ( $C^{1''}$ ). Found, %: C 67.89; H 6.32; N 4.72.  $[M]^+$  301.  $C_{17}H_{19}NO_4$ . Calculated, %: C 67.76; H 6.36; N 4.65.  $M$  301.34.

**3-Acyl-6-methyl-1,2,3,4-tetrahydropyridine-2,4-diones Ve and Vf (general procedure).** Triethylamine, 18.2 ml (0.13 mol), was added under stirring to a mixture of 0.1 mol of diketone **IIIa** or **IIIc** in 100 ml of methylene chloride, and a solution of 0.11 mol of pentanoyl or acetyl chloride in methylene chloride (1:1 by volume) was added dropwise over a period of 4 h under stirring. The mixture was then stirred for 0.5 h at

room temperature, and 28 ml (0.20 mol) of triethylamine and 0.5 ml (0.005 mol) of 2-hydroxy-2-methylpropanenitrile were added. As in the reactions with enol esters **IV**, the C–O isomerization was complete in 48 h (TLC). The mixture was treated as described above.

**1,6-Dimethyl-3-pentanoyl-1,2,3,4-tetrahydropyridine-2,4-dione (Ve).** Yield 73%. Oily substance. IR spectrum,  $\nu$ ,  $\text{cm}^{-1}$ : 1660 (C=O, lactam), 1620 (BuC=O), 1565 (C=C, enol).  $^1\text{H}$  NMR spectrum,  $\delta$ , ppm: 0.95 t (3H,  $\text{CH}_3\text{CH}_2\text{CH}_2$ ,  $J = 7.3$  Hz), 1.41 m (2H,  $\text{CH}_3\text{CH}_2\text{CH}_2$ ), 1.65 m (2H,  $\text{CH}_2\text{CH}_2\text{CH}_2$ ), 2.35 s (3H, 6- $\text{CH}_3$ ), 3.17 t (2H,  $\text{CH}_2\text{CH}_2\text{CO}$ ,  $J = 7.7$  Hz), 3.45 s (3H,  $\text{NCH}_3$ ), 5.84 s (1H, 5-H), 15.67 s (1H, OH). Found, %: C 64.69; H 7.65; N 6.40.  $[M]^+$  223.  $\text{C}_{12}\text{H}_{17}\text{NO}_3$ . Calculated, %: C 64.55; H 7.67; N 6.27.  $M$  223.27.

**3-Acetyl-1-benzyl-6-methyl-1,2,3,4-tetrahydropyridine-2,4-dione (Vf).** Yield 75%. Oily substance. IR spectrum,  $\nu$ ,  $\text{cm}^{-1}$ : 1660 (C=O, lactam), 1620 (MeC=O), 1575 (C=C, enol).  $^1\text{H}$  NMR spectrum,  $\delta$ , ppm: 2.28 s (3H, 6- $\text{CH}_3$ ), 2.74 s (3H,  $\text{CH}_3\text{CO}$ ), 5.27 s (2H,  $\text{CH}_2\text{C}_6\text{H}_5$ ), 5.86 s (1H, 5-H), 7.13 d (2H,  $\text{H}_{\text{arom}}$ ), 7.27 m (1H,  $\text{H}_{\text{arom}}$ ), 7.33 m (2H,  $\text{H}_{\text{arom}}$ ), 15.65 s (1H, OH). Found, %: C 70.13; H 5.78; N 5.42.  $[M]^+$  257.  $\text{C}_{15}\text{H}_{15}\text{NO}_3$ . Calculated, %: C 70.02; H 5.88; N 5.44.  $M$  257.19.

**Enamines VIa–VIh (general procedure).** Triketone **Va**, **Vb**, **Vd**, or **Ve**, 5 mmol, was dissolved in 50 ml of toluene, 6.5 mmol of the corresponding amine was added, and the mixture was heated under reflux for 4–6 (aliphatic amines) or 60 h (aromatic amines), the progress of the reaction being monitored by TLC. The mixture was washed with 5% hydrochloric acid and water, the aqueous phases were combined and extracted with chloroform (3  $\times$  10 ml), the chloroform extracts were combined with the toluene layer, dried over anhydrous magnesium sulfate, and passed through a thin layer of silica gel, and the solvent was evaporated under reduced pressure.

**1,6-Dimethyl-3-[1-(phenylamino)ethylidene]-1,2,3,4-tetrahydropyridine-2,4-dione (VIa).** Yield 52%, mp 116–118°C (from diethyl ether). IR spectrum,  $\nu$ ,  $\text{cm}^{-1}$ : 3060, 1660, 1620, 1560.  $^1\text{H}$  NMR spectrum,  $\delta$ , ppm: 2.25 s (3H, 6- $\text{CH}_3$ ), 2.71 s (3H,  $\text{CH}_3\text{CN}$ ), 3.42 s (3H,  $\text{CH}_3\text{N}$ ), 5.79 s (1H, 5-H), 7.10–7.40 m (5H,  $\text{C}_6\text{H}_5$ ), 17.1 s (1H, NH). Found, %: C 70.39; H 6.21; N 11.02.  $[M]^+$  256.  $\text{C}_{15}\text{H}_{16}\text{N}_2\text{O}_2$ . Calculated, %: C 70.29; H 6.29; N 10.93.  $M$  256.30.

**3-[1-(2-Allylamino)ethylidene]-1,6-dimethyl-1,2,3,4-tetrahydropyridine-2,4-dione (VIb).** Yield

85%, mp 69–69.5°C (from diethyl ether). IR spectrum,  $\nu$ ,  $\text{cm}^{-1}$ : 3070, 1660, 1620, 1600, 1560.  $^1\text{H}$  NMR spectrum,  $\delta$ , ppm: 2.22 s (3H, 6- $\text{CH}_3$ ), 2.70 s (2H,  $\text{CH}_3\text{CN}$ ), 3.35 s (3H,  $\text{CH}_3\text{N}$ ), 4.13 t (2H,  $\text{NHCH}_2$ ,  $J = 5.5$  Hz), 5.29 m (2H,  $\text{CH}_2=\text{CH}$ ), 5.70 s (1H, 5-H), 5.90 m (1H,  $\text{CH}_2=\text{CH}$ ), 15.3 s (1H, NH). Found, %: C 65.61; H 7.42; N 12.66.  $[M]^+$  220.  $\text{C}_{12}\text{H}_{16}\text{N}_2\text{O}_2$ . Calculated, %: C 65.43; H 7.32; N 12.72.  $M$  220.27.

**3-[1-(2-Allylamino)pentylidene]-1,6-dimethyl-1,2,3,4-tetrahydropyridine-2,4-dione (VIc).** Yield 87%, mp 38–40°C (from diethyl ether). IR spectrum,  $\nu$ ,  $\text{cm}^{-1}$ : 3070, 1655, 1620, 1590, 1560.  $^1\text{H}$  NMR spectrum,  $\delta$ , ppm: 0.97 t (3H,  $\text{CH}_3\text{CH}_2$ ,  $J = 7.4$  Hz), 1.51 m (2H,  $\text{CH}_2\text{CH}_3$ ), 2.22 s (3H, 6- $\text{CH}_3$ ), 3.13 m (2H,  $\text{CH}_2\text{CH}_2\text{CH}_2$ ), 3.35 s (3H,  $\text{CH}_3\text{N}$ ), 4.13 t (2H,  $\text{NHCH}_2$ ,  $J = 5.5$  Hz), 5.30 m (2H,  $\text{CH}_2=\text{CH}$ ), 5.67 s (1H, 5-H), 5.90 m (1H,  $\text{CH}_2=\text{CH}$ ), 15.3 s (1H, NH). Found, %: C 68.77; H 8.55; N 10.46.  $[M]^+$  262.  $\text{C}_{15}\text{H}_{22}\text{N}_2\text{O}_2$ . Calculated, %: C 68.67; H 8.45; N 10.68.  $M$  262.35.

**3-{1-[(4-Methoxyphenyl)amino]propylidene}-6-methyl-1-propyl-1,2,3,4-tetrahydropyridine-2,4-dione (VI d).** Yield 44%, mp 89–90°C (from diethyl ether). IR spectrum,  $\nu$ ,  $\text{cm}^{-1}$ : 3080, 1670, 1620, 1590, 1560.  $^1\text{H}$  NMR spectrum,  $\delta$ , ppm: 1.00 t (3H,  $\text{CH}_3\text{CH}_2\text{CH}_2$ ,  $J = 7.5$  Hz), 1.22 t (3H,  $\text{NHCCH}_2\text{CH}_3$ ,  $J = 7.5$  Hz), 1.68 m (2H,  $\text{CH}_3\text{CH}_2\text{CH}_2$ ), 2.30 s (3H, 6- $\text{CH}_3$ ), 3.10 q (2H,  $\text{NHCCH}_2\text{CH}_3$ ,  $J = 7.5$  Hz), 3.80 m (2H,  $\text{NCH}_2$ ), 3.84 s (3H,  $\text{OCH}_3$ ), 5.73 s (1H, 5-H), 6.94 d (2H,  $\text{C}_6\text{H}_4$ ,  $J = 9.0$  Hz), 7.12 d (2H,  $\text{C}_6\text{H}_4$ ,  $J = 9.0$  Hz), 16.85 s (1H, NH). Found, %: C 69.77; H 7.61; N 8.37.  $[M]^+$  328.  $\text{C}_{19}\text{H}_{24}\text{N}_2\text{O}_3$ . Calculated, %: C 69.49; H 7.37; N 8.53.  $M$  328.41.

**3-[1-(Benzylamino)propylidene]-6-methyl-1-propyl-1,2,3,4-tetrahydropyridine-2,4-dione (VIe).** Yield 82%. Oily substance. IR spectrum,  $\nu$ ,  $\text{cm}^{-1}$ : 3080, 1670, 1620, 1590, 1560.  $^1\text{H}$ ,  $\delta$ , ppm: 0.95 t (3H,  $\text{CH}_3\text{CH}_2\text{CH}_2$ ,  $J = 7.5$  Hz), 1.23 t (3H,  $\text{NHCCH}_2\text{CH}_3$ ,  $J = 7.5$  Hz), 1.68 m (2H,  $\text{CH}_3\text{CH}_2\text{CH}_2$ ), 2.23 s (3H, 6- $\text{CH}_3$ ), 3.24 q (2H,  $\text{NHCCH}_2\text{CH}_3$ ,  $J = 7.5$  Hz), 3.82 m (2H,  $\text{NCH}_2$ ), 4.78 d (2H,  $\text{NHCH}_2\text{Ph}$ ,  $J = 5.5$  Hz), 5.68 s (1H, 5-H), 7.20–7.42 m (5H,  $\text{C}_6\text{H}_5$ ), 15.65 s (1H, NH). Found, %: C 73.10; H 7.65; N 9.01.  $[M]^+$  312.  $\text{C}_{19}\text{H}_{24}\text{N}_2\text{O}_2$ . Calculated, %: C 73.05; H 7.74; N 8.97.  $M$  312.41.

**3-[1-(Benzylamino)butylidene]-6-methyl-1-(4-methoxyphenyl)-1,2,3,4-tetrahydropyridine-2,4-dione (VI f).** Yield 85%, mp 154–155°C (from ethyl acetate). IR spectrum,  $\nu$ ,  $\text{cm}^{-1}$ : 3080, 1670, 1630, 1590, 1560.  $^1\text{H}$  NMR spectrum,  $\delta$ , ppm: 1.00 t (3H,  $\text{CH}_3\text{CH}_2\text{CH}_2$ ,  $J = 7.4$  Hz), 1.64 m (4H,  $\text{CH}_3\text{CH}_2\text{CH}_2$ ),

1.82 s (3H, 6-CH<sub>3</sub>), 3.13 t (1.6H,  $J = 8.0$  Hz) and 3.31 br.s (0.4H, NHCH<sub>2</sub>CH<sub>2</sub>), 3.83 s (3H, OCH<sub>3</sub>), 4.64 br.s (0.4H) and 4.71 d (1.6H,  $J = 5.5$  Hz) (NHCH<sub>2</sub>Ph), 5.74 s (1H, 5-H), 6.97 m (2H, H<sub>arom</sub>), 7.08 m (2H, H<sub>arom</sub>), 7.33–7.38 m (5H, H<sub>arom</sub>), 15.59 s (1H, NH). Found, %: C 73.98; H 6.65; N 7.03.  $[M]^+$  390. C<sub>24</sub>H<sub>26</sub>N<sub>2</sub>O<sub>3</sub>. Calculated, %: C 73.82; H 6.71; N 7.17. *M* 390.48.

**1-(4-Methoxyphenyl)-6-methyl-3-[1-(4-tolylamino)butylidene]-1,2,3,4-tetrahydropyridine-2,4-dione (VIg).** Yield 44%, mp 150–152°C (from diethyl ether). IR spectrum,  $\nu$ , cm<sup>-1</sup>: 3060, 1660, 1625, 1560, 1520. <sup>1</sup>H NMR spectrum,  $\delta$ , ppm: 0.80 t (2.4H,  $J = 7.4$  Hz) and 0.89–0.98 m (0.6H) (CH<sub>3</sub>CH<sub>2</sub>CH<sub>2</sub>), 1.60 m (2H, CH<sub>3</sub>CH<sub>2</sub>CH<sub>2</sub>), 1.86 s (3H, 6-CH<sub>3</sub>), 2.39 s (CH<sub>3</sub>C<sub>6</sub>H<sub>4</sub>), 2.98 t (1.6H,  $J = 7.7$  Hz) and 3.14 m (0.4H) (CH<sub>2</sub>CH<sub>2</sub>CH<sub>3</sub>), 3.84 s (3H, CH<sub>3</sub>O), 5.81 s (1H, 5-H), 6.97–7.25 m (8H, H<sub>arom</sub>), 14.7 s (0.2H) and 16.85 s (0.8H) (NH). <sup>13</sup>C NMR spectrum,  $\delta_c$ , ppm: 14.60 (C<sup>4''</sup>), 20.86 (C<sup>3''</sup>), 21.81 (6-Me), 31.95 (C<sup>2''</sup>), 47.20 (CH<sub>2</sub>Ph), 55.44 (MeO), 101.92 (C<sup>3</sup>), 107.50 (C<sup>5</sup>), 114.76 (C<sup>3'</sup>), 127.16 (C<sup>2'''</sup>), 127.96 (C<sup>4'''</sup>), 129.03 (C<sup>3'''</sup>), 129.76 (C<sup>2'</sup>), 131.87 (C<sup>1'</sup>), 136.04 (C<sup>1'''</sup>), 149.12 (C<sup>6</sup>), 159.17 (C<sup>4'</sup>), 165.03 (C<sup>2</sup>), 183.73 (C<sup>4</sup>), 179.37 (C<sup>1''</sup>). Found, %: C 74.01; H 6.61; N 7.30.  $[M]^+$  390. C<sub>24</sub>H<sub>26</sub>N<sub>2</sub>O<sub>3</sub>. Calculated, %: C 73.82; H 6.71; N 7.17. *M* 390.48.

**1,6-Dimethyl-3-[1-(4-methoxyphenylamino)pentylidene]-1,2,3,4-tetrahydropyridine-2,4-dione (VIh).** Yield 48%, mp 120–121°C (from methanol). IR spectrum,  $\nu$ , cm<sup>-1</sup>: 3050, 1650, 1625, 1570, 1555. <sup>1</sup>H NMR spectrum,  $\delta$ , ppm: 0.79 t (3H, CH<sub>3</sub>CH<sub>2</sub>CH<sub>2</sub>,  $J = 7.1$  Hz), 1.29 m (2H, CH<sub>3</sub>CH<sub>2</sub>CH<sub>2</sub>), 1.56 m (2H, CH<sub>3</sub>CH<sub>2</sub>CH<sub>2</sub>), 2.26 s (3H, 6-CH<sub>3</sub>), 3.05 m (2H, NH-CCH<sub>2</sub>CH<sub>2</sub>), 3.80 s (3H, NCH<sub>3</sub>), 3.84 s (3H, CH<sub>3</sub>O), 5.74 s (1H, 5-H), 6.93 m (2H, H<sub>arom</sub>), 7.08 m (2H, H<sub>arom</sub>), 16.85 s (1H, NH). Found, %: C 69.33; H 7.17; N 8.47.  $[M]^+$  328. C<sub>19</sub>H<sub>24</sub>N<sub>2</sub>O<sub>3</sub>. Calculated, %: C 69.49; H 7.37; N 8.53. *M* 328.41.

**Enol ethers VIIa and VIIb (general procedure).** Triketone **Vd** or **Ve**, 0.05 mol, was dissolved in 50 ml of toluene, 2.8 g (0.02 mol) of finely powdered anhydrous potassium carbonate and 0.57 ml (0.06 mol) of dimethyl sulfate were added, and the mixture was heated for 6–8 h under reflux with stirring. The mixture was cooled and filtered from potassium carbonate, the precipitate was washed on a filter with toluene, the filtrate was passed through a thin layer of silica gel, the solvent was removed under reduced pressure, and the residue was recrystallized from a mixture of diethyl ether and petroleum ether.

**4-Methoxy-1,6-dimethyl-3-pentanoyl-1,2-dihydropyridin-2-one (VIIa).** Yield 98%, mp 52–53°C. IR spectrum,  $\nu$ , cm<sup>-1</sup>: 1700, 1655, 1590, 1560. <sup>1</sup>H NMR spectrum,  $\delta$ , ppm: 0.90 t (3H, CH<sub>3</sub>CH<sub>2</sub>CH<sub>2</sub>,  $J = 7.4$  Hz), 1.35 m (2H, CH<sub>3</sub>CH<sub>2</sub>CH<sub>2</sub>), 1.64 m (2H, CH<sub>2</sub>-CH<sub>2</sub>CH<sub>2</sub>), 2.39 s (3H, 6-CH<sub>3</sub>), 2.83 t (2H, CH<sub>2</sub>CH<sub>2</sub>CO,  $J = 7.7$  Hz), 3.48 s (3H, NCH<sub>3</sub>), 3.82 s (3H, OCH<sub>3</sub>), 5.95 s (1H, 5-H). <sup>13</sup>C NMR spectrum,  $\delta_c$ , ppm: 13.97 (C<sup>5''</sup>), 21.74 (6-Me), 22.41 (C<sup>4''</sup>), 26.11 (C<sup>3''</sup>), 30.82 (C<sup>1'</sup>), 43.54 (C<sup>2''</sup>), 56.02 (MeO), 94.84 (C<sup>5</sup>), 112.64 (C<sup>3</sup>), 127.16 (C<sup>2'''</sup>), 149.25 (C<sup>6</sup>), 162.03 (C<sup>2</sup>), 164.08 (C<sup>4</sup>), 203.18 (C<sup>1''</sup>). Found, %: C 65.64; H 7.95; N 6.02.  $[M]^+$  237. C<sub>13</sub>H<sub>19</sub>NO<sub>3</sub>. Calculated, %: C 65.80; H 8.07; N 5.90. *M* 237.30.

**3-Butanoyl-4-methoxy-1-(4-methoxyphenyl)-6-methyl-1,2-dihydropyridin-2-one (VIIb).** Yield 97%, mp 142–143°C. IR spectrum,  $\nu$ , cm<sup>-1</sup>: 1710, 1650, 1610, 1585. <sup>1</sup>H NMR spectrum,  $\delta$ , ppm: 0.92 t (3H, CH<sub>3</sub>CH<sub>2</sub>CH<sub>2</sub>,  $J = 7.4$  Hz), 1.67 m (2H, CH<sub>3</sub>CH<sub>2</sub>CH<sub>2</sub>), 2.03 s (3H, 6-CH<sub>3</sub>), 2.85 t (2H, CH<sub>2</sub>CH<sub>2</sub>CO,  $J = 7.4$  Hz), 3.84 s (3H, NCH<sub>3</sub>), 3.88 s (3H, OCH<sub>3</sub>), 6.02 s (1H, 5-H), 7.08 m (2H, H<sub>arom</sub>), 7.14 m (2H, H<sub>arom</sub>). <sup>13</sup>C NMR spectrum,  $\delta_c$ , ppm: 13.93 (C<sup>4''</sup>), 17.58 (C<sup>3''</sup>), 22.50 (6-Me), 45.75 (C<sup>2''</sup>), 55.67 (4'-MeO), 56.36 (4-MeO), 94.95 (C<sup>5</sup>), 113.05 (C<sup>3</sup>), 115.18 (C<sup>3'</sup>), 129.06 (C<sup>2'</sup>), 130.70 (C<sup>1'</sup>), 150.09 (C<sup>6</sup>), 159.81 (C<sup>4</sup>), 162.84 (C<sup>2</sup>), 165.30 (C<sup>4</sup>), 202.96 (C<sup>1''</sup>). Found, %: C 65.64; H 7.95; N 6.02.  $[M]^+$  315. C<sub>18</sub>H<sub>21</sub>NO<sub>4</sub>. Calculated, %: C 68.55; H 6.71; N 4.44. *M* 315.37.

**Enamines VIIa–VIIId (general procedure).** A mixture of 0.01 mol of 4-methoxy-pyridine **VIIa** or **VIIb** and 0.013 mmol of the corresponding amine in 50 ml of toluene was heated for 18–24 h under reflux (TLC). The mixture was then treated as described above for enamino derivatives **VI**.

**4-Benzylamino-3-butanoyl-1-(4-methoxyphenyl)-6-methyl-1,2-dihydropyridin-2-one (VIIa).** Yield 85%, mp 145–146°C (from methanol). IR spectrum,  $\nu$ , cm<sup>-1</sup>: 3070, 1660, 1630, 1560, 1520. <sup>1</sup>H NMR spectrum,  $\delta$ , ppm: 0.92 t (3H, CH<sub>3</sub>CH<sub>2</sub>CH<sub>2</sub>,  $J = 7.4$  Hz), 1.63 m (2H, CH<sub>3</sub>CH<sub>2</sub>CH<sub>2</sub>), 1.87 s (3H, 6-CH<sub>3</sub>), 3.10 t (2H, CH<sub>2</sub>CH<sub>2</sub>CO,  $J = 7.4$  Hz), 3.83 s (3H, OCH<sub>3</sub>), 4.52 d (2H, NHCH<sub>2</sub>,  $J = 5.8$  Hz), 5.73 s (1H, 5-H), 6.99 m (2H, H<sub>arom</sub>), 7.07 m (2H, H<sub>arom</sub>), 7.26–7.40 m (5H, H<sub>arom</sub>), 11.53 t (1H, NH,  $J = 5.45$  Hz). <sup>13</sup>C NMR spectrum,  $\delta_c$ , ppm: 14.12 (C<sup>4''</sup>), 18.15 (C<sup>3''</sup>), 22.58 (6-Me), 46.31 (C<sup>2''</sup>), 46.64 (CH<sub>2</sub>Ph), 55.56 (4'-MeO), 94.48 (C<sup>5</sup>), 101.73 (C<sup>3</sup>), 114.95 (C<sup>3'</sup>), 127.05 (C<sup>2'''</sup>, C<sup>6'''</sup>), 127.66 (C<sup>4'''</sup>), 128.98 (C<sup>3'''</sup>, C<sup>5'''</sup>), 129.46 (C<sup>2'</sup>), 131.48 (C<sup>1'</sup>), 150.80 (C<sup>6</sup>), 137.41 (C<sup>1'''</sup>), 159.45 (C<sup>4</sup>), 164.52 (C<sup>2</sup>), 159.82 (C<sup>4</sup>), 204.82 (C<sup>1''</sup>).

Found, %: C 73.78; H 6.60; N 7.25.  $[M]^+$  390.  $C_{24}H_{26}N_2O_3$ . Calculated, %: C 73.82; H 6.71; N 7.17.  $M$  390.48.

**3-Butanoyl-1-(4-methoxyphenyl)-6-methyl-4-(4-tolylamino)-1,2-dihydropyridin-2-one (VIIIb).** Yield 43%, mp 153–154°C (from diethyl ether). IR spectrum,  $\nu$ ,  $cm^{-1}$ : 3070, 1665, 1630, 1560, 1520.  $^1H$  NMR spectrum,  $\delta$ , ppm: 0.95 t (3H,  $CH_3CH_2CH_2$ ,  $J = 7.4$  Hz), 1.67 m (2H,  $CH_3CH_2$ ), 1.83 s (3H,  $CH_3C_6H_4$ ), 2.38 s (3H, 6- $CH_3$ ), 3.15 t (2H,  $CH_2CH_2CO$ ,  $J = 7.7$  Hz), 3.84 s (3H,  $NCH_3$ ), 5.87 s (1H, 5-H), 7.01 m (2H,  $H_{arom}$ ), 7.09 m (2H,  $H_{arom}$ ), 7.14 m (2H,  $H_{arom}$ ), 7.21 m (2H,  $H_{arom}$ ), 12.63 s (1H, NH). Found, %: C 73.98; H 6.65; N 7.03.  $[M]^+$  390.  $C_{24}H_{26}N_2O_3$ . Calculated, %: C 73.82; H 6.71; N 7.17.  $M$  390.48.

**4-(4-Methoxyphenylamino)-1,6-dimethyl-3-pentanoyl-1,2-dihydropyridin-2-one (VIIIc).** Yield 45%, mp 98–99°C (from hexane). IR spectrum,  $\nu$ ,  $cm^{-1}$ : 3050, 1650, 1620, 1565, 1520.  $^1H$  NMR spectrum,  $\delta$ , ppm: 0.96 t (3H,  $CH_3CH_2CH_2$ ,  $J = 7.4$  Hz), 1.42 m (2H,  $CH_3CH_2CH_2$ ), 1.67 m (2H,  $CH_3CH_2CH_2$ ), 2.19 s (3H, 6- $CH_3$ ), 3.20 t (2H,  $CH_2CH_2CO$ ,  $J = 7.4$  Hz), 3.41 s (3H,  $NCH_3$ ), 3.83 s (3H,  $OCH_3$ ), 5.69 s (1H, 5-H), 6.91 m (2H,  $H_{arom}$ ), 7.10 m (2H,  $H_{arom}$ ), 12.26 s (1H, NH). Found, %: C 69.48; H 7.27; N 8.35.  $[M]^+$  328.  $C_{19}H_{24}N_2O_3$ . Calculated, %: C 69.49; H 7.37; N 8.53.  $M$  328.41.

**4-Allylamino-1,6-dimethyl-3-pentanoyl-1,2-dihydropyridin-2-one (VIIId).** Yield 65%, mp 59–60°C (from diethyl ether). IR spectrum,  $\nu$ ,  $cm^{-1}$ : 3090, 1660, 1620, 1570, 1520.  $^1H$  NMR spectrum,  $\delta$ , ppm: 0.94 t (3H,  $CH_3CH_2CH_2$ ,  $J = 7.4$  Hz), 1.39 m (2H,  $CH_3CH_2CH_2$ ), 1.63 m (2H,  $CH_3CH_2CH_2$ ), 2.29 s (3H, 6- $CH_3$ ), 3.14 t (2H,  $CH_2CH_2CO$ ,  $J = 7.69$  Hz), 3.41 s (3H,  $NCH_3$ ), 3.88 m (2H,  $NHCH_2$ ), 5.23 m (2H,  $CH_2=CH$ ), 5.62 s (1H, 5-H), 5.88 m (1H,  $CH_2=CH$ ), 10.90 s (1H, NH).  $^{13}C$  NMR spectrum,  $\delta_C$ , ppm: 14.30 ( $C^5$ ), 22.03 (6-Me), 22.83 ( $C^4$ ), 27.33 ( $C^3$ ), 30.52 ( $C^1$ ), 44.10 ( $C^2$ ), 44.85 ( $C^{1''}$ ), 94.56 ( $C^5$ ), 101.74 ( $C^3$ ), 133.33 ( $C^{2''}$ ), 116.89 ( $C^{3''}$ ), 150.15 ( $C^6$ ), 164.06 ( $C^2$ ), 159.01 ( $C^4$ ), 205.01 ( $C^{1''}$ ). Found, %: C 68.77; H 8.55; N 10.46.  $[M]^+$  262.  $C_{15}H_{22}N_2O_2$ . Calculated, %: C 68.67; H 8.45; N 10.68.  $M$  262.35.

## REFERENCES

- O'Hagan, D., *Nat. Prod. Rep.*, 2000, vol. 17, p. 435.
- Malle, E., Stadlbauer, W., Ostermann, G., Hofmann, B., Leis, H.J., and Kostner, G.M., *Eur. J. Med. Chem.*, 1990, vol. 25, p. 137.
- Wat, C.-K., McInnes, A.G., Smith, D.G., Wright, J.L.C., and Vining, L.C., *Can. J. Chem.*, 1977, vol. 55, p. 4090.
- Schmidt, K., Riese, U., Li, Z., and Hamburger, M., *J. Nat. Prod.*, 2003, vol. 66, p. 378.
- Irlapati, N.G., Adlington, R.M., Conte, A., Pritchard, G.J., Marquez, R., and Baldwin, J.E., *Tetrahedron*, 2004, vol. 60, p. 9307.
- Hirono, Y., Ishikawa, H., Iwataki, I., and Sawaki, M., JPN Patent Appl. no. 75-154272, 1975; *Chem. Abstr.*, 1976, vol. 85, no. 78010s.
- Grina, J., FRG Patent Appl. no. 3820538, 1989; *Chem. Abstr.*, 1989, vol. 110, no. 212625x.
- Curtis, J.K., EPV Patent Appl. no. 249149, 1987; *Chem. Abstr.*, 1988, vol. 109, no. 73334r.
- Geach, N.J., Gilmour, J., Hatton, L.R., and Smith, P.H.G., EPV Patent Appl. no. 278742, 1988; *Chem. Abstr.*, 1989, vol. 110, no. 38898b.
- Kappe, T. and Schnell, B., *J. Heterocycl. Chem.*, 1996, vol. 33, p. 663.
- Schnell, B. and Kappe, T., *Monatsh. Chem.*, 1998, vol. 129, p. 871.
- Alonso, P., Martin-Leon, N., Quinteiro, M., and Soto, J.L., *Justus Liebigs Ann. Chem.*, 1990, p. 841.
- Takahashi, S., Kakinuma, N., Uchida, K., Hashimoto, R., Yanagishima, T., and Nakagawa, A., *J. Antibiot.*, 1998, vol. 51, p. 596.
- Kappe, T. and Kappe, C.O., *J. Heterocycl. Chem.*, 1989, vol. 26, p. 1555.
- Rubinov, D.B., Zheldakova, T.A., and Rubinova, I.L., *Russ. J. Org. Chem.*, 2004, vol. 40, p. 1329.
- Rubinov, D.B., Rubinova, I.L., and Akhrem, A.A., *Russ. J. Org. Chem.*, 1995, vol. 31, p. 386.
- Rubinov, D.B., Rubinova, I.L., and Akhrem, A.A., *Chem. Rev.*, 1999, vol. 99, p. 1047.
- Budnikova, M.V. and Rubinov, D.B., *Russ. J. Org. Chem.*, 2001, vol. 37, p. 1478.
- Budnikova, M.V., Gulyakevich, O.V., Zheldakova, T.A., Mikhal'chuk, A.L., and Rubinov, D.B., *Russ. J. Org. Chem.*, 2002, vol. 38, p. 1696.
- Zheldakova, T.A., Budnikova, M.V., Rubinova, I.L., and Rubinov, D.B., *Russ. J. Org. Chem.*, 2003, vol. 39, p. 1772.
- Gulyakevich, O.V., Mikhal'chuk, A.L., Verenich, A.I., Rubinov, D.B., Zenyuk, A.A., and Akhrem, A.A., *Enaminy v organicheskom sinteze* (Enamines in Organic Synthesis), Yekaterinburg: Ural. Otd. Ross. Akad. Nauk, 1996, p. 111.
- Budnikova, M.V., Zheldakova, T.A., Rubinov, D.B., and Mikhal'chuk, A.L., *Russ. J. Org. Chem.*, 2001, vol. 37, p. 293.
- Budnikova, M.V., Rubinov, D.B., and Mikhal'chuk, A.L., *Khim. Geterotsykl. Soedin.*, 2002, p. 1067.