# A Simple Synthesis of Some New Thienopyridine and Thienopyrimidine Derivatives 

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#### Abstract

A number of pyridothienopyridine (2, 3), pyridothienopyrimidine (6, 10), pyridothienotriazine (13), pyrimidothienopyrimidine (15, 16, 17a,b) and thienoimidazo-triazines (18) were obtained via interaction of 3-amino-5-phenylamino-2,4-dicarbonitrile (1) with different reagents.


Keywords: pyridothienopyridines, pyridothienopyrimidines, thienotriazopyrimidines, pyridothienotriazines, pyrimidothienopyrimidines.

## Introduction

Many thienopyridines have been evaluated pharmacologically and have been fund to show activity against, for example, diabetes mellitus [1-3], as analgesics and antiinflammatories [4-6], sedatives [4], anticoagulants [6], antiartherosclerotics [7], and as gonadotropin releasing hormone antagonists [8]. Moreover the pyridothienopyrimidines showed analgesic and antiinflammatory activity [9].

In earlier communications, we have reported the syntheses and reactions of various thiophenes and fused thiophenes [10-12], which were prepared in one-pot reactions using phase transfer techniques (PTC). Also, several papers have reported the synthesis of polyfused heterocycles containing different nuclei [13-16], starting from fused thiophenes prepared in our laboratory.

## Results and Discussion

We report herein the synthesis of some new polyfunctionally substituted thienopyridines, thienopyrimidines and pyridothienopyrimidines starting with 3-amino-5-phenylamino-2,4-dicarbonitrile (1). Compound 1 was prepared in one-pot reaction using PTC conditions $\left[\mathrm{K}_{2} \mathrm{CO}_{3} /\right.$ benzene/ tetrabutylammonium bromide (TBAB)] from malononitrile, PhNCS and $\mathrm{ClCH}_{2} \mathrm{CN}$ compounds in a 1:1:1 molar ratio, (Scheme 1) [12].

## Scheme 1



Compound 1 was allowed to react with two moles of ethyl cyanoacetate or diethyl malonate in the presence of ammonium acetate-acetic acid at $200^{\circ} \mathrm{C}$ [17] to give the corresponding 4,9-diamino-2,7-dioxo-6-phenyl-1,2,6,7-tetrahydropyrido[2',3':4,5]thieno[2,3-b]pyridin-3,8-dicarbonitrile (2) or 4,9-diamino-3,8-diethoxycarbonyl-6-phenyl-1,2,6,7-tetrahydro[2',3':4,5]thieno[2,3-b]pyridin-2,7-dione (3).

## Scheme 2



Reaction of compound 1 with formamide gave the corresponding 4 amino-6-phenylamino-thieno[3,2-d]pyrimidin-7-carbonitrile (4) which was allowed then allowed to react with benzaldehyde to yield 6 phenylamino-4-benzalaminothieno[3,2-d]pyrimidin-7-carbonitrile (5). 9-Amino-7-oxo-6-phenyl-4-phenylmethanimido-6,7-dihydropyrido-[3',2':4,5]thieno[3,2-d]pyrimidin-8-carbonitrile (6) was obtained by treating compound $\mathbf{5}$ with ethyl cyanoacetate (Scheme 2 ).

The condensation of compound $\mathbf{1}$ with triethyl orthoformate in refluxing acetic anhydride gave ethyl N -(2,4-dicyano-5-phenylamino-3-yl)metanimidate (7) which underwent further cyclization upon treatment with hydrazine hydrate at room temperature affording 3-amino-3,4-dihydro-4-imino-6-phenylaminothieno[3,2-d]pyrimidin-7-carbonitrile (8) (Scheme 3).

## Scheme 3



Reaction of compound $\mathbf{8}$ with triethyl orthoformate gave 8-phenylaminothieno[3,2-d][1,2,4]-triazolo[3,2-flpyrimidin-7-carbonitrile (9). Fusion of compound 9 with ethyl cyanoacetate in the presence of ammonium acetate yielded 7-amino-9-oxo-10-phenyl-9,10-dihydropyrido[3',2':4,5]-thieno[3,2-d][1,2,4]triazolo[3,2-flpyrimidin-8-carbonitrile (10). Diazotisation of compound 1 with $\mathrm{NaNO}_{2} /$ concentrated HCl led to the formation of 4-chloro-6-phenylaminothieno[3,2-d]-1,2,3-triazin-7-carbonitrile (11) which, in turn, was allowed to react with ethyl cyanoacetate to give 9-amino-4-chloro-7-oxo-6-phenyl-6,7dihydropyrido $\left[3^{\prime}, 2^{\prime}: 4,5\right]$ thieno[3,2-d][1,2,3]triazin-8-carbonitrile (12). Compound 12 was further treated with hydrazine hydrate to afford 9-amino-4-hydrazino-7-oxo-6-phenyl-6,7-dihydropyrido-[3',2':4,5]-thieno[3,2-d][1,2,3]triazin-8-carbonitrile (13).

Incorporation of the imidazolyl moiety in pyrimidothienopyrimidine, imidazothienopyrimidine and imidazothienotriazine systems were achieved by converting the nitrile group of compound $\mathbf{1}$ into a dihydroimidazolyl residue, followed by some additional reactions [18]. Thus, the interaction of compound 1 with ethylenediamine in the presence of carbon disulfide afforded 3 -amino-2,4-di(4,5-dihydro- $1 \mathrm{H}-2$ -imidazolyl)-5-phenylaminothiophene (14) (Scheme 4).

## Scheme 4



Treatment of compound $\mathbf{1 4}$ with triethyl orthoformate, benzaldehyde, dimethythiomethylenemalononitrile or dimethylthio-2-acetylbuten-3-one furnished 5,6,12-triphenyl-2,3,5,6,9,10-hexahydroimidazo[1,2c]imidazo[ $\left.2^{\prime \prime}, 1^{\prime \prime}: 6^{\prime}, 1^{\prime}\right]$ pyrimido[4',5':4,5]thieno[3,2-e]pyrimidine (15), 5-ethoxy-6-phenyl-2,3,5,6,-9,10-hexahydroimidazo[1,2-c]imidazo[2", $\left.1^{\prime \prime}: 6^{\prime}, 1^{\prime}\right]$ pyrimido[4',5':4,5]-thieno[3,2-e]-pyrimidine (16), 2-(12-(1,1-dicyanomethylidene)-6-phenyl-2,3,5,6,9,10,12,13-octahydro-imidazo[1,2-c]imidazo[2",1":6',1']pyrimido-[4',5':4,5]thieno[3,2-e]pyrimidin-5-yliden)malononitrile (17a) or 2-(12-(1-acetyl-2-oxopropylidene)-6-phenyl-2,3,5,6,9,10,12,13-octahydroimidazo[1,2-c]imidazo[2",1":6',1']-pyrimido[4',5':4,5]thieno[3,2-
e]pyrimidin-5-yliden)-2,4-pentandione (17b), respectively. In addition 7 -(4,5-dihydro-1H-2-imidazolyl)-2,3-dihydro-8-phenylaminoimidazo[1,2-c]-thieno[2,3-e][1,2,3]-triazine (18) was obtained through the treatment of compound 14 with nitrous acid (Scheme 5). Analytical and spectral data of the newly synthesized compounds were in agreement with the proposed structures (Tables 1 and 2).

## Scheme 5



## Experimental

## General

All melting points were determined on a Koffler melting points apparatus and are uncorrected. IR spectra were obtained on a Nicolet 710 FT-IR spectrometer. ${ }^{1} \mathrm{H}$-NMR spectra were recorded on a Varian EM 360 A at 60 MHz using TMS as an internal reference. Elemental analyses were carried out with an elemental
analyzer model 240 C. Satisfactory microanalysis ( $\mathrm{C} \pm 0.4, \mathrm{H} \pm 0.4, \mathrm{~N} \pm 0.3 \%$ ) were obtained for all newly prepared compounds.

4,9-diamino-2,7-dioxo-6-phenyl-1,2,6,7-tetrahydropyrido[2',3':4,5]thieno[2,3-b]pyridin-3,8-dicarbonitrile (2) or 4,9-diamino-3,8-diethoxycarbonyl-6-phenyl-1,2,6,7-tetrahydro[2',3':4,5]-thieno[2,3-blpyridin-2,7-dione (3). A mixture of $\mathbf{1}(0.01 \mathrm{~mol})$, ethyl cyanoacetate or diethyl malonate ( 0.02 mol ), ammonium acetate ( 6 g ) and acetic acid ( 1.2 mL ) was heated with stirring at $200^{\circ} \mathrm{C}$ for 2 h , left to cool and then triturated with ethanol. The solid product thus formed was collected by filtration. Compound $\mathbf{2}$ (from ethyl cyanoacetate) was recrystallized from ethanol as yellow needles; m. p. $200{ }^{\circ}$ C. Compound $\mathbf{3}$ (from diethyl malonate) was recrystallized from ethanol as orange crystals; m. p. $>320^{\circ} \mathrm{C}$.

4-Amino-6-phenylaminothieno[3,2-d]pyrimidin-7-carbonitrile (4). A solution of $\mathbf{1}$ ( 0.001 mol ) in formamide ( 10 mL ) was refluxed for 2 h . The precipitate that formed on cooling was filtered off and recrystallized from ethanol to give white powder, m.p. $>271{ }^{\circ} \mathrm{C}$.

6-Phenylamino-4-benzalaminothieno[3,2-d]pyrimidin-7-carbonitrile (5). An equimolar mixture of 4 $(0.01 \mathrm{~mol})$ and benzaldehyde ( 0.01 mol ) was dissolved in $\mathrm{EtOH}(30 \mathrm{~mL})$ in the presence of a few drops of piperidine. The reaction mixture was refluxed for 2 h and left to cool. The solid product was filtered off and recrystallized from pyridine as white crystals; m.p. $243^{\circ} \mathrm{C}$.

General Procedure for the Preparation of Thienopyridone Derivatives 6, 10 or 12. A mixture of 1 ( 0.01 $\mathrm{mol})$, ethyl cyanoacetate ( 0.01 mol ), ammonium acetate $(3 \mathrm{~g})$ and acetic acid $(0.6 \mathrm{~mL})$ was heated with stirring at $200^{\circ} \mathrm{C}$ for 2 h , then left to cool and triturated with ethanol. The solid product, so formed, was collected by filtration and recrystallized from appropriate solvents.

9-Amino-7-oxo-6-phenyl-4-phenylmethanimido-6,7-dihydropyrido[3', $\left.2^{\prime}: 4,5\right]$ thieno[3,2-d]-pyrimidin-8carbonitrile (6). Recrystallized from pyridine as white crystals; m.p. $220^{\circ} \mathrm{C}$.

7-Amino-9-oxo-10-phenyl-9, 10-dihydropyrido[3',2':4,5]thieno[3,2-d][1,2,4]triazolo-[3,2-f]-pyrimidin-8-carbonitrile (10). Recrystallized from acetonitrile as yellow crystals; m.p. $236^{\circ} \mathrm{C}$.

9-Amino-4-chloro-7-oxo-6-phenyl-6,7-dihydropyrido-[3', $\left.2^{\prime}: 4,5\right]$ thieno[3,2-d][1,2,3]triazin-8-carbonitrile (12). Recrystallized from acetonitrile as yellow crystals; m.p. 158-160 ${ }^{\circ} \mathrm{C}$.

Ethyl N-(2,4-dicyano-5-phenylamino-3-yl)metanimidate (7). A mixture of 1 ( 0.005 mol ), triethyl orthoformate ( 3 mL ) and acetic anhydride ( 20 mL ) was heated under reflux for 5 h . After cooling the precipitated solid was filtered off and recrystallized from ethanol as white crystals; m.p. $215^{\circ} \mathrm{C}$.

3-Amino-3,4-dihydro-4-imino-6-phenylaminoyhieno[3,2-d]pyrimidin-7-carbonitrile (8). Hydrazine hydrate $(80 \%)(4 \mathrm{~mL})$ was added to a suspension of $7(0.005 \mathrm{~mol})$ in dioxane $(40 \mathrm{~mL})$. The reaction mixture was stirred at room temperature for 1 h . The precipitate which formed was filtered off, washed with water, dried in air and recrystallized from dioxane as white crystals; m.p. $185^{\circ} \mathrm{C}$.

8-Phenylaminothieno[3,2-d][ 1,2,4]triazolo[3,2-f]pyrimidin-7-carbonitrile (9). Compound 8 (0.001 mol ) in an excess of triethyl orthoformate ( 7 mL ) was refluxed for 1 h . After cooling, the precipitated product was collected by filtration and recrystallized from ethanol-chloroform mixture as white needles; m.p. $299^{\circ} \mathrm{C}$.

4-Chloro-6-phenylamino-thieno[3,2-d]-1,2,3-triazin-7-carbonitrile (11). A solution of ( 0.01 mol ) sodium nitrite in 10 mL of water was added to a cold solution of $\mathbf{1}(0.005 \mathrm{~mol})$ in acetic acid ( 30 mL ) and concentrated hydrochloric acid ( 15 mL ). After completion of the addition, the ice bath was removed and stirring continued for an additional 2 h . The crude product obtained was recrystallized from ethanol as white needles; m.p. $201^{\circ} \mathrm{C}$.

9-Amino-4-hydrazino-7-oxo-6-phenyl-6,7-dihydropyrido-[3',2':4,5]thieno[3,2-d][1,2,3]triazin-8-
carbonitrile (13). A mixture of $\mathbf{1 2}(0.002 \mathrm{~mol})$ and hydrazine hydrate ( 3 mL ) in ethanol ( 20 mL ) was refluxed for 1 h . The precipitate that separated after cooling was recrystallized from dioxane as white crystals; m.p. $262^{\circ} \mathrm{C}$.

3-Amino-2,4-di(4,5-dihydro-1H-2-imidazolyl)-5-phenylaminothiophene (14). To a suspension of 1 ( 0.002 mol ), ethylenediamine ( 3 mL ) and carbon disulfide ( 1 mL ) were added dropwise. The reaction mixture was heated on a water bath for 2 h . The precipitated solid was triturated with ethanol ( 10 mL ), filtered off and recrystallized from ethanol to give golden yellow crystals; m.p. $197^{\circ} \mathrm{C}$.

## 5,6,12-Triphenyl-2,3,5,6,9,10-hexahydroimidazo[1,2-c]imidazo-[2",1":6',1']pyrimido[4',5':4,5]-

thieno-[3,2-e]pyrimidine (15). A mixture of $\mathbf{1 4}(0.005 \mathrm{~mol})$, benzaldehyde ( 0.01 mol ) and acetic acid ( 15 mL ) was heated under reflux for 5 h . The precipitated solid was collected and recrystallized from dioxane in the form of white needles; m.p. $215^{\circ} \mathrm{C}$.

5-Ethoxy-6-phenyl-2,3,5,6,-9,10-hexahydroimidazo[1,2-c]imidazo[2", $\left.1^{\prime \prime}: 6^{\prime}, 1^{\prime}\right]$ pyrimido[4',5':4,5]-thieno[3,2-e]pyrimidine (16). Compound $\mathbf{1 4}(0.001 \mathrm{~mol})$ in triethyl orthoformate $(14 \mathrm{~mL})$ was heated under reflux for 3 h . The precipitated solid was collected and recrystallized from pyridine as pale yellow crystals; m.p. $163^{\circ} \mathrm{C}$.

Preparation of 2-(12-(1,1-dicyanomethylidene)-6-phenyl-2,3,5,6,9,10,12,13-octahydroimidazo[1,2c]imidazo[ $\left.2^{\prime \prime}, 1^{\prime \prime}: 6^{\prime}, 1^{\prime}\right]$ pyrimido[4',5':4,5]thieno[3,2-e]pyrimidin-5-yliden)malononitrile (17a) and 2-(12-(1-Acetyl-2-oxopropylidene)-6-phenyl-2,3,5,6,9,10,12,13-octahydroimidazo[1,2-c]imidazo-[2", $1^{\prime \prime}$ : 6', 1']pyrimido[4',5':4,5]thieno[3,2-e]pyrimidin-5-yliden)-2,4-pentandione (17b). A mixture of 1 (0.001
mol ) and dimethylthiomethylenemalononitrile or 2-acetyl-1,1-dimethylthiobuten-3-one ( 0.002 mol ) in ethanol $(20 \mathrm{~mL})$ was heated under reflux for 24 h . The separated solids were collected and recrystallized from the appropriate solvent. Compound 17a: gray crystals; m.p. $172{ }^{\circ} \mathrm{C}$ (from dioxane). Compound 17b: orange crystals; m.p. $179{ }^{\circ} \mathrm{C}$ (from methanol).

7-(4,5-Dihydro-1H-2-imidazolyl)-2,3-dihydro-8-phenylaminoimidazo[1,2-c]thieno[2,3-e][1,2,3]-
triazine (18). To a solution of $\mathbf{1}(0.001 \mathrm{~mol})$ in concentrated sulphuric acid ( 2 mL ) and glacial acetic acid ( 10 mL ), sodium nitrite ( 0.003 mol ) dissolved in 5 mL water was added dropwise with constant stirring during 10 minutes. The mixture was stirred without heating for additional 1 h and then diluted with water. The forming precipitate was filtered off and recrystallized from ethanol as white needles; m.p. $269^{\circ} \mathrm{C}$.

Table 1: Analytical data for the newly prepared compounds

| Product | Yield <br> $\boldsymbol{\%}$ | Mol.Form. <br> Mol. Wt. | Analysis(\%) Calcd./Found |  |  |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: |
|  |  |  | $\boldsymbol{H}$ | $\boldsymbol{N}$ | $\boldsymbol{S}$ |  |
| $\mathbf{2}$ | 76 |  | 57.75 | 2.67 | 22.44 | 8.56 |
|  |  |  | 57.64 | 2.73 | 22.55 | 8.62 |
| $\mathbf{3}$ | 81 | $\mathrm{C}_{22} \mathrm{H}_{20} \mathrm{~N}_{4} \mathrm{O}_{6} \mathrm{~S}$ | 56.40 | 4.27 | 17.93 | 6.83 |
|  |  | 468.48 | 56.48 | 4.05 | 17.85 | 6.77 |
| $\mathbf{4}$ | 92 | $\mathrm{C}_{13} \mathrm{H}_{9} \mathrm{~N}_{5} \mathrm{~S}$ | 58.41 | 3.63 | 26.28 | 11.97 |
|  |  | 267.30 | 58.56 | 3.55 | 25.99 | 11.88 |
| $\mathbf{5}$ | 85 | $\mathrm{C}_{20} \mathrm{H}_{13} \mathrm{~N}_{5} \mathrm{~S}$ | 67.58 | 3.66 | 19.69 | 9.00 |
|  |  | 355.41 | 67.67 | 3.77 | 19.58 | 8.98 |
| $\mathbf{6}$ | 77 | $\mathrm{C}_{23} \mathrm{H}_{14} \mathrm{~N}_{6} \mathrm{OS}$ | 65.38 | 3.31 | 19.88 | 7.57 |
|  |  | 422.46 | 65.44 | 3.23 | 19.92 | 7.61 |
| $\mathbf{7}$ | 89 | $\mathrm{C}_{15} \mathrm{H}_{12} \mathrm{~N}_{4} \mathrm{OS}$ | 60.82 | 4.05 | 18.90 | 10.82 |
|  |  | 269.21 | 61.02 | 3.98 | 19.03 | 1076 |
| $\mathbf{8}$ | 90 | $\mathrm{C}_{13} \mathrm{H}_{10} \mathrm{~N}_{6} \mathrm{~S}$ | 55.30 | 3.54 | 29.75 | 11.33 |
|  |  | 282.32 | 55.22 | 3.46 | 29.66 | 11.43 |
| $\mathbf{9}$ | 78 | $\mathrm{C}_{14} \mathrm{H}_{8} \mathrm{~N}_{6} \mathrm{~S}$ | 57.52 | 2.74 | 28.74 | 11.94 |
|  |  | 292.31 | 57.33 | 2.87 | 28.65 | 11.99 |
| $\mathbf{1 0}$ | 66 | $\mathrm{C}_{17} \mathrm{H}_{9} \mathrm{~N}_{7} \mathrm{OS}$ | 56.81 | 2.50 | 27.27 | 8.90 |
|  |  | 359.36 | 56.77 | 2.56 | 27.30 | 9.12 |
| $\mathbf{1 1}$ | 75 | $\mathrm{C}_{12} \mathrm{H}_{6} \mathrm{~N}_{5} \mathrm{SCl}$ | 50.10 | 2.09 | 24.33 | 11.12 |
|  |  | 287.72 | 49.99 | 1.98 | 24.15 | 10.98 |


| $\mathbf{1 2}$ | 81 | $\mathrm{C}_{15} \mathrm{H}_{7} \mathrm{~N}_{6} \mathrm{OSC}$ | 50.77 | 1.97 | 23.68 | 9.02 |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: |
|  |  | 354.77 | 50.85 | 2.01 | 23.44 | 8.94 |
| $\mathbf{1 3}$ | 64 | $\mathrm{C}_{15} \mathrm{H}_{10} \mathrm{~N}_{8} \mathrm{OS}$ | 51.42 | 2.85 | 31.97 | 9.13 |
|  |  | 350.35 | 51.61 | 2.82 | 31.99 | 8.99 |
| $\mathbf{1 4}$ | 91 | $\mathrm{C}_{16} \mathrm{H}_{18} \mathrm{~N}_{6} \mathrm{~S}$ | 58.87 | 5.61 | 25.73 | 9.80 |
|  |  | 326.41 | 58.84 | 5.54 | 25.62 | 9.91 |
| $\mathbf{1 5}$ | 89 | $\mathrm{C}_{30} \mathrm{H}_{24} \mathrm{~N}_{6} \mathrm{~S}$ | 71.97 | 4.79 | 211.40 | 6.39 |
|  |  | 500.62 | 71.89 | 4.81 | 21.32 | 6.41 |
| $\mathbf{1 6}$ | 78 | $\mathrm{C}_{20} \mathrm{H}_{20} \mathrm{~N}_{6} \mathrm{OS}$ | 61.20 | 5.10 | 21.40 | 8.15 |
|  |  | 392.47 | 60.32 | 4.94 | 21.22 | 8.00 |
| $\mathbf{1 7 a}$ | 85 | $\mathrm{C}_{24} \mathrm{H}_{14} \mathrm{~N}_{10} \mathrm{~S}$ | 60.75 | 2.95 | 22.50 | 6.74 |
|  |  | 474.50 | 60.56 | 3.02 | 22.65 | 6.61 |
| $\mathbf{1 7 b}$ | 82 | $\mathrm{C}_{28} \mathrm{H}_{26} \mathrm{~N}_{6} \mathrm{O}_{4} \mathrm{~S}$ | 61.97 | 4.79 | 15.48 | 5.89 |
|  |  | 542.61 | 62.13 | 4.81 | 15.50 | 5.75 |
| $\mathbf{1 8}$ | 89 | $\mathrm{C}_{16} \mathrm{H}_{15} \mathrm{~N}_{7} \mathrm{~S}$ | 56.95 | 4.44 | 29.04 | 9.48 |
|  |  | 337.40 | 57.02 | 4.32 | 30.12 | 9.51 |

Table 2: Spectroscopic data of the newly prepared compounds.

| Product | IR ( $\mathbf{K B r})^{\text {b }}$ |  |
| :---: | :---: | :---: |
| 2 | $3430,3330,3300\left(2 \mathrm{NH}_{2}\right), 3180(\mathrm{NH})$, <br> 2222, 2210 ( 2 CN ), 1640, 1630 (C=O) | $\begin{aligned} & \begin{array}{l} 8.20 \quad(\mathrm{~s}, 1 \mathrm{H}, \mathrm{NH}), \\ \left(\mathrm{br}, 2 \mathrm{H}, \mathrm{NH}_{2}\right) \end{array} \end{aligned}$ |
| 3 | $\begin{aligned} & 3420,3300,3250\left(2 \mathrm{NH}_{2}\right), \quad 3190(\mathrm{NH}), \\ & 1720,1710(\mathrm{C}=\mathrm{O} \text { ester), } 1630(\mathrm{C}=\mathrm{O}) . \end{aligned}$ | 8.30-7.80 (m,5H,arom.), 6.30-6.10 (br,2H, $\mathrm{NH}_{2}$ ), 4.80- <br> $4.60\left(\mathrm{br}, 2 \mathrm{H}, \mathrm{NH}_{2}\right), 2.80\left(\mathrm{~s}, 3 \mathrm{H}, \mathrm{COCH}_{3}\right), 2.10\left(\mathrm{~s}, 3 \mathrm{H}, \mathrm{CH}_{3}\right)$. |
| 4 | $3340,3300,\left(\mathrm{NH}_{2}\right), 3200(\mathrm{NH}), 2218$ (CN). | 9.10 ( $\mathrm{s}, 1 \mathrm{H}, \mathrm{NH}$ ) , 8.40 ( $\mathrm{s}, 1 \mathrm{H}, \mathrm{CH}-$ Pyridine) , $\quad 7.70-7.20$ (m, 5 H, arom.), $5.50\left(\mathrm{~s}, 2 \mathrm{H}, \mathrm{NH}_{2}\right)$. |
| 5 | $3200(\mathrm{NH}), 2210$ (CN). | $9.10 \quad(\mathrm{~s}, 1 \mathrm{H}, \mathrm{NH}), \quad 8.40 \quad(\mathrm{~s}, 1 \mathrm{H}, \mathrm{CH}-$ Pyridine $), \quad 7.90-7.30$ (m,10H,arom.), 5.50-5.10 (br,2H, $\mathrm{NH}_{2}$ ). |
| 6 | $\begin{aligned} & 3350,3240 \quad\left(\mathrm{NH}_{2}\right), 2220(\mathrm{CN}), 1640 \\ & (\mathrm{C}=\mathrm{O}) . \end{aligned}$ | 8.40 ( $\mathrm{s}, 1 \mathrm{H}, \mathrm{CH}$-Pyridine), 8.10 ( $\mathrm{s}, 1 \mathrm{H}, \mathrm{N}=\mathrm{CH}$ ), 7.90-7.00 (m,10H,arom.), $6.20\left(\mathrm{~s}, 2 \mathrm{H}, \mathrm{NH}_{2}\right)$. |
| 7 | $\begin{array}{\|l} \hline 3180(\mathrm{NH}), 2220,2210 \quad(\mathrm{CN}), 1630 \\ (\mathrm{C}=\mathrm{N}) . \end{array}$ | $\begin{array}{\|l} 8.50(\mathrm{~s}, 1 \mathrm{H}, \mathrm{NH}), 7.70(\mathrm{~s}, 1 \mathrm{H}, \mathrm{CH}), 7.70-7.00(\mathrm{~m}, 5 \mathrm{H}, \text { arom. }), \\ 3.80-3.40\left(\mathrm{q}, 2 \mathrm{H}, \mathrm{OCH}_{2}\right), 1.25-1.00\left(\mathrm{t}, 3 \mathrm{H}, \mathrm{CH}_{3}\right) . \end{array}$ |
| 8 | $\begin{array}{\|l} \hline 3430,3330\left(\mathrm{NH}_{2}\right), 3200,3180(2 \mathrm{NH}), \\ 2190(\mathrm{CN}) . \end{array}$ | $\begin{array}{llll} \begin{array}{l} 8.20-8.00 \end{array} \quad(\mathrm{br}, 2 \mathrm{H}, 2 \mathrm{NH}), \quad 7.80 \quad(\mathrm{~s}, 1 \mathrm{H}, \mathrm{CH}), \quad 7.70-7.20 \\ (\mathrm{~m}, 5 \mathrm{H}, \text { arom. }), 5.20\left(\mathrm{~s}, 2 \mathrm{H}, \mathrm{NH}_{2}\right) . \end{array}$ |
| 9 | $\begin{aligned} & \begin{array}{l} 3180 \quad(\mathrm{NH}), \quad 2220 \quad(\mathrm{CN}), \quad 1600 \\ (\mathrm{C}=\mathrm{N}) . \end{array} \\ & \hline \end{aligned}$ | 8.80 ( $\mathrm{s}, 1 \mathrm{H}, \mathrm{NH}$ ), 8.40 ( $\mathrm{s}, 1 \mathrm{H}, \mathrm{CH}$-Pyridine), 8.10 ( $\mathrm{s}, 1 \mathrm{H}, \mathrm{CH}-$ triazole), $7.50-7.00$ (m,5H,arom.), $6.20\left(\mathrm{~s}, 2 \mathrm{H}, \mathrm{NH}_{2}\right)$. |


| 10 | 3400, 3330, $3300 \quad\left(2 \mathrm{NH}_{2}\right), \quad 2217$ (CN), 1640 (C=O), $1600(\mathrm{C}=\mathrm{N})$. | $\begin{aligned} & 8.50(\mathrm{~s}, 1 \mathrm{H}, \mathrm{CH} \text {-Pyridine }), 8.20(\mathrm{~s}, 1 \mathrm{H}, \mathrm{CH} \text {-triazole }), 7.50- \\ & 7.00(\mathrm{~m}, 5 \mathrm{H}, \text { arom. }), 6.20\left(\mathrm{~s}, 2 \mathrm{H}, \mathrm{NH}_{2}\right) \text {. } \end{aligned}$ |
| :---: | :---: | :---: |
| 11 | 3320 (NH), 2210 (CN), 1640 (C=N). | 8.40 (s,1H,NH), 7.70-7.20 (m,5H,arom.). |
| 12 | $\begin{aligned} & 3340,3300,3270\left(2 \mathrm{NH}_{2}\right), 2220(\mathrm{CN}), \\ & 1640(\mathrm{C}=\mathrm{O}), 1620(\mathrm{C}=\mathrm{N}) . \end{aligned}$ | 7.70-7.00 (m,5H,arom.), 6.20-6.00 (br, $2 \mathrm{H},\left(\mathrm{NH}_{2}\right)$. |
| 13 | $\begin{array}{lrl} 3350,3330,3210 & \left(2 \mathrm{NH}_{2}\right), & 3180, \\ (\mathrm{NH}), 2220(\mathrm{CN}), 1640(\mathrm{C}=\mathrm{O}) & \end{array}$ | $\begin{aligned} & 8.00(\mathrm{~s}, 1 \mathrm{H}, \mathrm{NH}), 7.80-7.20(\mathrm{~m}, 5 \mathrm{H}, \mathrm{arom} .), 6.10\left(\mathrm{~s}, 2 \mathrm{H}, \mathrm{NH}_{2}\right), \\ & 5.00\left(\mathrm{~s}, 2 \mathrm{H}, \mathrm{NH}_{2}\right) . \end{aligned}$ |
| 14 | $\begin{aligned} & 3390,3260 \quad\left(\mathrm{NH}_{2}\right), \quad 3200,3180,3100 \\ & (3 \mathrm{NH}) . \end{aligned}$ | $\begin{aligned} & 8.40(\mathrm{~s}, 1 \mathrm{H}, \mathrm{NH}), 8.10(\mathrm{~s}, 1 \mathrm{H}, \mathrm{NH}), 7.70-7.10(\mathrm{~m}, 5 \mathrm{H}, \text { arom. }) \text {, } \\ & 5.60\left(\mathrm{~s}, 2 \mathrm{H}, \mathrm{NH}_{2}\right), 4.00-3.50\left(\mathrm{~m}, 8 \mathrm{H}, 4 \mathrm{CH}_{2}\right. \text {-imidazole). } \end{aligned}$ |
| 15 | 3080 (CH-arom.), 1640 (C=N) | $\begin{array}{\|l\|ll} \hline 7.90-7.10 \quad(\mathrm{~m}, 15 \mathrm{H}, \text { arom. }), \quad 4.00-3.40 \quad\left(\mathrm{~m}, 9 \mathrm{H}, \quad 4 \mathrm{CH}_{2}-\right. \\ \text { imidazole }+1 \mathrm{H}, \mathrm{CH}-\mathrm{Ph}) \end{array}$ |
| 16 | 3050 ( CH -arom.), 1640 ( $\mathrm{C}=\mathrm{N}$ ). | $\begin{aligned} & 9.10 \quad(\mathrm{~s}, 1 \mathrm{H}, \mathrm{CH}), \quad 7.70-7.20 \quad(\mathrm{~m}, 5 \mathrm{H}, \text { arom. }), \quad 4.00-3.40 \\ & \left(\mathrm{~m}, 10 \mathrm{H}, 4 \mathrm{CH}_{2} \text {-imidazole+ } \mathrm{OCH}_{2}\right), 1.25-1.00\left(\mathrm{t}, 3 \mathrm{H}, \mathrm{CH}_{3}\right) . \end{aligned}$ |
| 17a | $\begin{aligned} & 3200(\mathrm{NH}), 2220,2210,2197(4 \mathrm{CN}), \\ & 1620(\mathrm{C}=\mathrm{N}) . \end{aligned}$ | $\begin{aligned} & 9.00 \quad(\mathrm{~s}, 1 \mathrm{H}, \mathrm{NH}), \quad 7.70-7.20 \quad(\mathrm{~m}, 5 \mathrm{H}, \text { arom. }), \quad 4.00-3.40 \\ & \left(\mathrm{~m}, 8 \mathrm{H}, 4 \mathrm{CH}_{2} \text {-imidazole }\right) . \end{aligned}$ |
| 17b | $\begin{aligned} & 3200(\mathrm{NH}), 1690,1670(\mathrm{C}=\mathrm{O}), 1640 \\ & (\mathrm{C}=\mathrm{N}) . \end{aligned}$ | $\begin{aligned} & 8.90 \quad(\mathrm{~s}, 1 \mathrm{H}, \mathrm{NH}), \quad 7.70-7.00 \quad(\mathrm{~m}, 5 \mathrm{H}, \text { arom. }), \quad 4.00-3.40 \\ & \left(\mathrm{~m},, 8 \mathrm{H}, 4 \mathrm{CH}_{2} \text {-imidazole }\right), 2.70-2.60\left(\mathrm{ss}, 12 \mathrm{H}, 2 \mathrm{COCH}_{3}\right) . \end{aligned}$ |
| 18 | 3050 ( CH -arom.), 1640 ( $\mathrm{C}=\mathrm{N}$ ). | $\begin{aligned} & 8.70(\mathrm{~s}, 1 \mathrm{H}, \mathrm{NH}), 8.40(\mathrm{~s}, 1 \mathrm{H}, \mathrm{NH}), 7.70-7.20(\mathrm{~m}, 5 \mathrm{H}, \text { arom. }) \text {, } \\ & 4.00-3.50\left(\mathrm{~m}, 8 \mathrm{H}, 4 \mathrm{CH}_{2}\right. \text {-imidazole). } \end{aligned}$ |

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