IR and $^1$H NMR spectral studies of some 2-Amino-4-isopropyl-6-methoxy-N-phenylpyrimidine-5-carboxamides: Assessment of substituent effects

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ABSTRACT

A series of 2-Amino-4-isopropyl-6-methoxy-N-phenylpyrimidine-5-carboxamide compounds have been synthesized from various substituted dithioacetal and guanidine. The purities of these compounds were checked by their physical constants, UV, IR, NMR and MASS spectral data. The IR and $^1$HNMR spectral data of these compounds have been correlated with Hammett sigma constants, F and R parameters using single and multi-linear regression analysis. From the results of statistical analysis, the effects of substituents on the spectral group frequencies have been discussed.

Keywords: 2-Amino-4-isopropyl-6-methoxy-N-phenylpyrimidine-5-carboxamides, IR spectra; NMR spectra; Substituent effects

1. INTRODUCTION

Heterocyclic chemistry comprises at least half of all organic chemistry research worldwide. In particular, heterocyclic structures form the basis of many pharmaceutical, agrochemical and medicinal products [1]. Nitrogen containing heterocyclic ring such as pyrimidine is a promising structural moiety for drug design. Pyrimidine derivatives form a component in a number of useful drugs and are associated with many biological and therapeutical activities [2]. Pyrimidine derivatives have been reported as anti-microbial [3], analgesic, anti-viral, anti-inflammatory [4], anti-HIV [5], anti-tubercular [6], anti-tumour [7], anti-malarial [8], cardiovascular [9] agents. Pyrimidine derivatives have been found to exhibit cytostatic [10-12], immune modulating [13,14] and antibacterial properties [15-21]. In particular, studies concerning the synthesis of pyrimido[4,5-d]-1,3-oxazines [22] showing considerable similarity to quinoline chemotherapeutic agents with an antibacterial effect. The spectral data of these 2-Amino-4-isopropyl-6-methoxy-N-phenyl pyrimidine-5-carboxamide compounds were useful for prediction of ground state equilibration like various conformers.
The effects of substituent on the infrared group frequencies (ν, cm\(^{-1}\)) and NMR chemical shifts (δ, ppm) of 2-amino-4-isopropyl-6-methoxy-N-phenylpyrimidine-5-carboxamide compounds. In their studies, they observed satisfactory and good correlations for each spectral data with Hammett substituent constants, F and R parameters.

2. EXPERIMENTAL

2.1 General

All chemicals used were purchased from Sigma-Aldrich and E-Merck chemical companies. Melting points of all chalcones were determined in open glass capillaries on Mettler FP51 melting point apparatus and are uncorrected. Infrared spectra (KBr, 4000-400 cm\(^{-1}\)) were recorded AVATAR-300 Fourier transform spectrophotometer. Bruker AV400 NMR spectrometer was used for recording NMR spectra operating at 400 MHz for \(^1\)H spectra and 125.46 MHz for \(^1^3\)C spectra in CDCl\(_3\) solvent using TMS as internal standard. Mass spectra of all chalcones were recorded on SHIMADZU spectrometer using chemical ionization technique.

2.2. Synthesis of substituted 2-amino-4-isopropyl-6-methoxy-N-phenylpyrimidine-5-carboxamides

All substituted 2-Amino-4-isopropyl-6-methoxy-N-phenylpyrimidine-5-carboxamide were synthesized and their purities were examined by procedure published in literature [23]. An equi-molar quantities of substituted dithioacetal (2 mmol) and guanidine (2 mmol), in the presence of sodium methoxide were refluxed for 6h and the completion of the reaction was monitored by thin layer chromatography.

Where X = H, o-Br, m-Cl, p-Cl, o-F, p-F, o-OCH\(_3\), p-OCH\(_3\), o-CH\(_3\), p-CH\(_3\)

Fig. 1. Substituted 2-Amino-4-isopropyl-6-methoxy-N-phenylpyrimidine-5-carboxamide
After completion of the reaction, the solid product was separated by filtration. Further the product was purified by column chromatography using dichloromethane and ethyl acetate mixture as eluent. The general structure of the carboxamides was shown in Fig. 1.

3. RESULTS AND DISCUSSION

In the present study the authors have studied the effects of substituents on the spectral data such as infrared spectral carbonyl and deformation modes (ν, cm⁻¹), the chemical shifts (δ, ppm) of –NH₂, -OCH₃ and –CONH protons of synthesized 2-amino-4-isopropyl-6-methoxy-N-phenylpyrimidine-5-carboxamide compounds with Hammett substituent constants, F and R parameters using single and multi-linear regression analysis for predicting the reactivity on the group frequencies.

Table 1. The infrared absorptions (ν, cm⁻¹) and NMR chemical shifts (δ, ppm) of substituted 2-amino-4-isopropyl-6-methoxy-N-phenylpyrimidine-5-carboxamide compounds.

| Entry | X   | IR (ν, cm⁻¹)   | NMR(δ, ppm) |
|-------|-----|---------------|-------------|
|       |     | C=O | C=N | NH₂ | -C-O-C- | NH₂ | OCH₃ | CONH |
| 1     | H   | 1618.34 | 1538.46 | 3414.34 | 1430.40 | 5.702 | 3.802 | 9.508 |
| 2     | o-Br| 1648.52 | 1586.02 | 3459.12 | 1261.08 | 5.608 | 3.808 | 9.600 |
| 3     | m-Cl| 1649.04 | 1588.44 | 3460.86 | 1246.48 | 5.640 | 3.806 | 9.602 |
| 4     | p-Cl| 1651.30 | 1588.86 | 3462.56 | 1248.50 | 5.602 | 3.810 | 9.582 |
| 5     | o-F | 1654.45 | 1590.68 | 3444.24 | 1260.56 | 5.624 | 3.820 | 9.586 |
| 6     | p-F | 1650.96 | 1588.46 | 3448.50 | 1262.84 | 5.612 | 3.904 | 9.624 |
| 7     | o-OCH₃| 1624.54 | 1508.34 | 3422.20 | 1151.46 | 5.694 | 3.782 | 9.504 |
| 8     | p-OCH₃| 1630.00 | 1509.21 | 3454.48 | 1052.24 | 5.584 | 3.764 | 9.488 |
| 9     | o-CH₃| 1620.98 | 1568.36 | 3440.27 | 1054.63 | 5.692 | 3.784 | 9.426 |
| 10    | p-CH₃| 1618.46 | 1538.40 | 3414.21 | 1060.61 | 5.576 | 3.688 | 9.482 |

3.1. IR spectral study

The synthesized compounds in the present study are shown in Fig. 1. The carbonyl C=N, NH₂ and -C-O-C- stretching frequencies (cm⁻¹) of substituted 2-amino-4-isopropyl-6-methoxy-N-phenylpyrimidine-5-carboxamides are tabulated in Table 1.

The infrared frequencies for carbonyl absorption are assigned based on the assignments made by Hays and Timmons [24] for CO at 1640, C=N at 1580, NH₂ at 3400 and -O-C-O- at 1200 cm⁻¹, respectively. The assigned infrared frequencies have been correlated with Hammett substituent constants and Swain-Lupton’s constants [25]. In this correlation the structure parameter Hammett equation employed is as shown in the following equation:
\[ \nu = \rho \sigma + \nu_0 \quad \ldots \text{(1)} \]

Where \( \nu \) is the carbonyl frequencies of substituted system and \( \nu_0 \) is the corresponding quantity of unsubstituted system; \( \sigma \) is a Hammett substituent constant, which in principle is characteristics of the substituent and \( \rho \) is a reaction constant which is depend upon the nature of the reaction.

The results of statistical analysis [26-40] were shown in Table 2. From Table 2, The Hammett substituent constants and F parameters correlated satisfactorily with CO stretches. The resonance component was fail in correlation. All the correlations have positive \( \rho \) values. This is due to the inability of prediction of the reactivity through the resonance on the carbonyl stretches and associated with the resonance-conjugative structure as shown in Fig. 2.

The Hammett substituent constants and F parameters gave satisfactory correlation with C=N stretches. The remaining Hammett substituent constant \( \sigma_R \) and R parameter have given poor correlation. This is due to the reasons stated in earlier and associated with the resonance conjugative structure. And all the compounds have positive \( \rho \) values.

Except R parameter, the Hammett substituent constants and F parameters gave satisfactory correlation with NH\(_2\) stretches. The reasons stated in earlier and associated with the resonance conjugative structure for the correlations fail with NH\(_2\) stretches.

The Hammett substituent constants and F and R parameters gave satisfactory correlation with –C-O-C- stretches and all correlations gave positive \( \rho \) values.

![Fig. 2. The resonance-conjugative structure.](image-url)
Table 2. Results of statistical analysis of infrared absorptions (ν, cm⁻¹) and NMR chemical shifts (δ, ppm) of substituted α-Amino-4-isopropyl-6-methoxy-N-phenylpyrimidine-5-carboxamide compounds with Hammett σ, σ⁺, σᵣ, σᵢ and F and R parameters.

| Frequency | Constants | r   | I     | ρ     | s     | n     | Correlated derivatives                                                                 |
|-----------|-----------|-----|-------|-------|-------|-------|----------------------------------------------------------------------------------------|
| νCO       | σ         | 0.907 | 1636.22 | 48.039 | 9.99  | 10    | H, o-Br, m-Cl, p-Cl, o-F, p-F, o-OCH₃, p-OCH₃, o-CH₃, p-CH₃                           |
|           | σ⁺        | 0.906 | 1637.34 | 24.045 | 11.82 | 10    | H, o-Br, m-Cl, p-Cl, o-F, p-F, o-OCH₃, p-OCH₃, o-CH₃, p-CH₃                           |
|           | σᵣ        | 0.832 | 1629.26 | -25.960 | 15.43 | 10    | H, o-Br, m-Cl, p-Cl, o-F, p-F, o-OCH₃, p-OCH₃, o-CH₃, p-CH₃                           |
|           | F         | 0.990 | 1619.56 | 50.717 | 6.85  | 10    | H, o-Br, m-Cl, p-Cl, o-F, p-F, o-OCH₃, p-OCH₃, o-CH₃, p-CH₃                           |
|           | R         | 0.883 | 1629.20 | -22.736 | 15.39 | 10    | H, o-Br, m-Cl, p-Cl, o-F, p-F, o-OCH₃, p-OCH₃, o-CH₃, p-CH₃                           |
| νC=N      | σ         | 0.988 | 1559.46 | 17.593 | 17.03 | 8     | H, o-Br, m-Cl, p-Cl, o-F, p-F, o-CH₃, p-CH₃                                         |
|           | σ⁺        | 0.983 | 1562.34 | 63.999 | 19.77 | 10    | H, o-Br, m-Cl, p-Cl, o-F, p-F, o-OCH₃, p-OCH₃, o-CH₃, p-CH₃                           |
|           | σᵣ        | 0.955 | 1536.58 | 81.973 | 29.90 | 8     | H, o-Br, m-Cl, p-Cl, o-F, p-F, o-CH₃, p-CH₃                                         |
|           | F         | 0.905 | 1536.87 | 70.158 | 29.48 | 8     | H, o-Br, m-Cl, p-Cl, o-F, p-F, o-CH₃, p-CH₃                                         |
|           | R         | 0.813 | 1567.25 | 20.525 | 35.57 | 10    | H, o-Br, m-Cl, p-Cl, o-F, p-F, o-CH₃, p-CH₃                                         |
| νNH₂      | σ         | 0.958 | 3441.68 | 43.701 | 16.21 | 10    | H, o-Br, m-Cl, p-Cl, o-F, p-F, o-OCH₃, p-OCH₃, o-CH₃, p-CH₃                           |
|           | σ⁺        | 0.938 | 3442.54 | 16.354 | 18.49 | 9     | H, o-Br, m-Cl, p-Cl, o-F, p-F, o-OCH₃, o-CH₃, p-CH₃                               |
|           | σᵣ        | 0.973 | 3424.30 | 60.861 | 15.53 | 10    | H, o-Br, m-Cl, p-Cl, o-F, p-F, o-OCH₃, p-OCH₃, o-CH₃, p-CH₃                           |
|     | $\sigma_R$ |  $\nu_{C\equiv O}$ |  $\delta N\equiv H$ |  $\delta O\equiv C_3$ | Alkyl Groups |
|-----|------------|---------------------|---------------------|----------------------|---------------|
| **F** | 0.959  | 3428.24  | 41.052  | 16.01  | 10  | H, o-Br, m-Cl, p-Cl, o-F, p-F, o-OCH$_3$, p-OCH$_3$, o-CH$_3$, p-CH$_3$ |
| **R** | 0.815  | 3437.17  | -14.956  | 16.69  | 10  | H, o-Br, m-Cl, p-Cl, o-F, p-F, o-OCH$_3$, p-OCH$_3$, o-CH$_3$, p-CH$_3$ |
| **σ** | 0.960  | 1200.27  | 289.665  | 103.14  | 9  | o-Br, m-Cl, p-Cl, o-F, p-F, o-OCH$_3$, p-OCH$_3$, o-CH$_3$, p-CH$_3$ |
| **σ’** | 0.962  | 1207.80  | 173.508  | 100.49  | 9  | Br, m-Cl, p-Cl, o-F, p-F, o-OCH$_3$, p-OCH$_3$, o-CH$_3$, p-CH$_3$ |
| **σ$_l$** | 0.931  | 1153.87  | 167.833  | 122.65  | 9  | Br, m-Cl, p-Cl, o-F, p-F, o-OCH$_3$, p-OCH$_3$, o-CH$_3$, p-CH$_3$ |
| **σ$_R$** | 0.921  | 1241.57  | 135.777  | 126.18  | 8  | H, o-Br, m-Cl, p-Cl, o-F, p-F, o-OCH$_3$, p-OCH$_3$, o-CH$_3$, p-CH$_3$ |
| **F** | 0.929  | 1158.87  | 130.579  | 123.47  | 9  | Br, m-Cl, p-Cl, o-F, p-F, o-OCH$_3$, p-OCH$_3$, o-CH$_3$, p-CH$_3$ |
| **R** | 0.926  | 1249.63  | 142.541  | 124.66  | 8  | H, o-Br, m-Cl, p-Cl, o-F, p-F, o-OCH$_3$, p-OCH$_3$, o-CH$_3$, p-CH$_3$ |
| **σ** | 0.921  | 5.633  | -0.404  | 0.04  | 7  | o-Br, m-Cl, p-Cl, o-F, p-F, o-OCH$_3$, p-CH$_3$ |
| **σ’** | 0.902  | 5.633  | -0.03  | 0.04  | 7  | o-Br, m-Cl, p-Cl, o-F, p-F, o-OCH$_3$, p-CH$_3$ |
| **σ$_l$** | 0.933  | 5.653  | -0.068  | 0.04  | 8  | H, o-Br, m-Cl, p-Cl, o-F, p-F, o-OCH$_3$, o-CH$_3$ |
| **σ$_R$** | 0.909  | 5.639  | 0.022  | 0.04  | 8  | H, o-Br, m-Cl, p-Cl, o-F, p-F, o-OCH$_3$, o-CH$_3$ |
| **F** | 0.935  | 0.653  | -0.060  | 0.04  | 8  | H, o-Br, m-Cl, p-Cl, o-F, p-F, o-OCH$_3$, o-CH$_3$ |
| **R** | 0.823  | 5.649  | 0.048  | 0.04  | 10  | H, o-Br, m-Cl, p-Cl, o-F, p-F, o-OCH$_3$, p-OCH$_3$, o-CH$_3$, p-CH$_3$ |
| **σ** | 0.946  | 3.795  | 0.097  | 0.05  | 8  | H, o-Br, m-Cl, p-Cl, o-F, p-F, o-OCH$_3$, p-OCH$_3$, o-CH$_3$, p-CH$_3$ |
| **σ’** | 0.940  | 3.798  | 0.048  | 0.05  | 8  | H, o-Br, m-Cl, p-Cl, o-F, p-F, o-OCH$_3$, o-CH$_3$, p-CH$_3$ |
| **σ$_l$** | 0.968  | 3.749  | 0.161  | 0.04  | 10  | H, o-Br, m-Cl, p-Cl, o-F, p-F, o-OCH$_3$, p-OCH$_3$, o-CH$_3$, p-CH$_3$ |
| **σ$_R$** | 0.935  | 3.769  | -0.097  | 0.05  | 10  | H, o-Br, m-Cl, p-Cl, o-F, p-F, o-OCH$_3$, p-OCH$_3$, o-CH$_3$, p-CH$_3$ |
| **F** | 0.972  | 3.749  | 0.140  | 0.03  | 10  | H, o-Br, m-Cl, p-Cl, o-F, p-F, o-OCH$_3$, p-OCH$_3$, o-CH$_3$, p-CH$_3$ |
Some of the single parameter correlations with Hammett substituent constants were not obeyed in the regression. While seeking these parameters in multi-regression, with F and R Swain-Lupton’s [25] constants, they gave satisfactory correlations with the infrared group frequencies. The multi correlation equations are given in (2-9).

\[
\nu_{\text{CO}}(\text{cm}^{-1}) = 1621.72(\pm 2.525) + 74.938(\pm 7.341)\sigma_i + 24.360(\pm 8.660)\sigma_R \quad \ldots(2)
\]

\[
\nu_{\text{CO}}(\text{cm}^{-1}) = 1624.31(\pm 2.271) + 68.336(\pm 6.205)F + 32.565(\pm 7.589)R \\
(R = 0.997, n = 10, P > 95 \%) \quad \ldots(3)
\]

\[
\nu_{\text{C} = \text{N}}(\text{cm}^{-1}) = 1554.07(\pm 14.189) + 140.567(\pm 41.252)\sigma_i + 121.410(\pm 48.659)\sigma_R \\
(R = 0.979, n = 10, P > 95 \%) \quad \ldots(4)
\]

\[
\nu_{\text{C} = \text{N}}(\text{cm}^{-1}) = 1556.91(\pm 10.225) + 144.556(\pm 27.933)F + 137.51(\pm 34.163)R \\
(R = 0.989, n = 10, P > 95 \%) \quad \ldots(5)
\]

\[
\nu_{\text{NH}_2}(\text{cm}^{-1}) = 3429.50(\pm 7.877) + 78.267(\pm 22.902)\sigma_i + 36.066(\pm 27.015)\sigma_R \\
(R = 0.979, n = 10, P > 95 \%) \quad \ldots(6)
\]

\[
\nu_{\text{NH}_2}(\text{cm}^{-1}) = 3432.97(\pm 9.430) + 58.631(\pm 25.762)F + 32.491(\pm 31.508)R \\
(R = 0.966, n = 10, P > 95 \%) \quad \ldots(7)
\]
\[ \nu C-O-C (cm^{-1}) = 1206.83(\pm 68.971) + 345.244(\pm 200.521)\sigma_I + 367.608(\pm 236.529)\sigma_R \quad \text{...(8)} \]
\[ \nu C-O-C (cm^{-1}) = 1223.19(\pm 59.862) + 369.467(\pm 163.537)F + 441.53(\pm 200.009)R \quad \text{...(9)} \]

3. 2. \(^1\)H NMR Spectral study

From the \(^1\)H NMR spectra of compounds of the chemical shifts (\(\delta\), ppm) \(\text{NH}_2\), -OCH\(_3\) and -CONH are assigned and tabulated in Table 2. These chemical shifts were correlated with Hammett substituent constants, F and R parameters. The statistical analysis [26-40] of these chemical shifts is presented in Table 2.

From Table 2, the chemical shifts values of –\(\text{NH}_2\) and –OCH\(_3\) (\(\delta\), ppm) correlated with Hammett substituent constants and F parameter gave satisfactory correlation except R parameter.

From Table 2, the chemical shifts values of –CONH (\(\delta\), ppm) correlated with Hammett substituent constants and F and R parameter gave satisfactory correlation. The other correlations were fail. This is due to the reasons stated earlier and associated with the resonance conjugative structure as shown in Fig. 2.

Some of the single parameter correlations with Hammett substituent constants were not obeyed in the regression. While seeking these parameters in multi-regression, with F and R Swain-Lupton’s [33] constants, they gave satisfactory correlations with the chemical shifts of (\(\delta\), ppm) H\(\alpha\) and H\(\beta\). The multi correlation equations are given in (10-15).

\[ \delta\text{NH}_2 \text{ (ppm)} = 5.648(\pm 0.030) - 0.085(\pm 0.088)\sigma_I - 0.035(\pm 0.103)\sigma_R \]
\[ (R = 0.935, n = 10, P > 90 \%) \quad \text{...(10)} \]

\[ \delta\text{NH}_2 \text{ (ppm)} = 5.653(\pm 0.029) - 0.060(\pm 0.080)F - 0.0003(\pm 0.098)R \]
\[ (R = 0.935, n = 10, P > 90 \%) \quad \text{...(11)} \]

\[ \delta\text{OCH}_3 \text{ (ppm)} = 3.752(\pm 0.026) + 0.009(\pm 0.038)\sigma_I + 0.152(\pm 0.073)\sigma_R \]
\[ (R = 0.969, n = 10, P > 95 \%) \quad \text{...(12)} \]

\[ \delta\text{OCH}_3 \text{ (ppm)} = 3.756(\pm 0.024) + 0.053(\pm 0.091)F + 0.164(\pm 0.064)R \]
\[ (R = 0.976, n = 10, P > 95 \%) \quad \text{...(13)} \]

\[ \delta\text{CONH} \text{ (ppm)} = 9.039(\pm 0.761) - 2.635(\pm 2.212)\sigma_I - 3.075(\pm 2.610)\sigma_R \]
\[ (R = 0.945, n = 10, P > 90 \%) \quad \text{...(14)} \]

\[ \delta\text{CONH} \text{ (ppm)} = 8.855(\pm 0.768) - 1.698(\pm 2.099)F - 2.630(\pm 2.567)R \]
\[ (R = 0.936, n = 10, P > 90 \%) \quad \text{...(15)} \]

4. CONCLUSIONS

Totally ten substituted 2-amino-4-isopropyl-6-methoxy-N-phenylpyrimidine-5-carboxamide compounds have been synthesized and their purities were examined by literature method. The IR and \(^1\)H NMR spectral data of these chalcones have been assigned and correlated with Hammett sigma constants, F and R parameters using single and multi-linear...
regression analysis. From the results of statistical analysis, the effects of substituents on the spectral group frequencies have been discussed. Satisfactory correlations coefficients (r) obtained for many single and all multi parameter correlations.

References

[1] T. Panneer Selvam, C. Richa James, P. Vijaysarathy Dniandev, S. Karyn Valzita, Res. in Phar. 2(4) (2012) 01-09.
[2] L. W. Hertel, G. B. Boder, J. S. Kroin, et al., Cancer Res. 50 (1990) 4417-4422.
[3] K. Desai, R. Patel, K. Chikhalia, J. Ind. Chem. 45(B) (2006) 773-778.
[4] A. E. Amr, M. S. Nermien, M. M. Abdulla, Monatsh. Chem., 138 (2007) 699-707.
[5] N. Fujiwara, T. Nakajima, Y. Ueda, H. Fujita, H. Kawakami, Bioorg. Med. Chem. 16 (2008) 9804-9816.
[6] L. Ballell, R. A. Field, G. A. C. Chung, R. J. Young, Bioorg. Med. Chem. Lett. 17 (2007) 1736-1740.
[7] E. Wagner, K. Al-Kadasi, M. Zimecki, W. Sawka-Dobrowolska, Eur. J. Med. Chem. 43 (2008) 2498-2504.
[8] K. Gorlitzer, S. Herbig, R. D. Walter, Pharmazie. 52 (1997) 670-672.
[9] W. ShenJeu, L. Cheng Tsung, T. Lo Ti, W. Jing Ru, K. Hrogn Huey. W. JihPyang, and L. C. Nan, Eur. J. Med. Chem. 40 (2005) 103-112.
[10] Z. Machon, J. Cieplik, Synthesis. 2 (1986) 142-144.
[11] Z. Machon, J. Cieplik, J. Pharmacol. Pharm. 40 (1988) 201-208.
[12] N. R. Mohamed, M. M. El-Saidi, Y. M. Ali, M. H. Elnagdi, Sci. Pharm. 75 (2007) 9-28.
[13] J. Cieplik, Z. MachoN, M. Zimecki, Z. Wieczorek, Arch. Immunol. Ther. Exp. 41(1993)11-15.
[14] J. Cieplik, Z. MachoN, M. Zimecki, Z. Wieczorek, Farmaco. 50 (1993) 131-132.
[15] J. Pluta, M. Flendrich, J. Cieplik, Boll. Chim. Farm. 135 (1996) 459-464.
[16] Y. A. Mostafa, M. A. Husein, A. A. Radwan, A. H. Kfary, Arch. Pharm. Res. Soc. Korea. 31 (2008) 279-293.
[17] G. T. Kim, R. J. Ryu, S. Lee Ch, H. Youn, Y. R. Cho, H. Y. Joo, J. Antibio. Jap. Res. (2004) 468-471.
[18] J. Cieplik, J. Pluta, A. Meler, Arch. Pharm. Pharm. Med. Chem. 330 (1997) 237-241.
[19] N. S. Habib, R. Soliman, A. A. El-Tombary, S. A. El-Hawash, O. G. Shaaban, Arch. Pharm. Res. Soc. Korea. 30 (2007) 1511-1520.
[20] J. Cieplik, J. Pluta, O. Gubrynowicz, Sci. Pharm. 68 (2000) 333-336.
[21] J. Cieplik, M. Raginia, J. Pluta, O. Gubrynowicz, I. Bryndal, T. Lis, Acta. Pol. Pharm. Drug. Res. 65 (2008) 427-434.
[22] Z. Macho, J. Cieplik, Eur. J. Med. Chem., Chim. Ther. 19 (1984) 359-363.
[23] H. R. Khunt, P. P. Pipaliya, S. M. Ghelani, J. S. Babariya, Y. T. Naliapara, *International Letters of Chemistry, Physics and Astronomy* 5 (2014) 134-121.

[24] W.P. Hays, C. J. Timmons., *Spectrochim. Acta.* 24A (1968) 323-334.

[25] C.G. Swain, E. C. Lupton Jr., *J. Am. Chem. Soc.* 90 (1968) 4328-4337.

[26] R. Ranganathan, R. Arulkumaran, D. Kamalakkannan, G. Vanangamudi, G. Thirunayraran. *IUP J. Chem.* 4(2) (2011) 60-70.

[27] K. Sathiyamoorthi, V. Mala, R. Suresh, S. P. Sakhthinathan, D. Kamalakkannan, K. Ranganathan, R. Arulkumaran, S. Vijayakumar, G. Vanangamudi, G. Thirunayraran, *International Letters of Chemistry, Physics and Astronomy* 7 (2013) 102-119.

[28] V. Mala, K. Sathiyamoorthi, S. P. Sakhthinathan, D. Kamalakkannan, R. Suresh, G. Vanangamudi, G. Thirunayraran, *Q. Science Connect.* (2013) 7, DOI: http://dx.doi.org/ 10.5339/ connect.2013.7

[29] S. P. Sakhthinathan, R. Suresh, V. Mala, K. Sathiyamoorthi, D. Kamalakkannan, K. Ranganathan, R. Arulkumaran, S. Vijayakumar, R. Sundararajan, G. Vanangamudi, G. Thirunayraran, *International Letters of Chemistry, Physics and Astronomy* 6 (2013) 77-90.

[30] R. Suresh, D. Kamalakkannan, K. Ranganathan, R. Arulkumaran, R. Sundararajan, S.P. Sakhthinathan, S. Vijayakumar, K. Sathiyamoorthi, V. Mala, G. Vanangamudi, K. Thirumurthy, P. Mayavel, G. Thirunayraran, *Spectrochim. Acta Part A.*, 101 (2013) 239-248.

[31] K. Sathiyamoorthi, V. Mala, S.P. Sakhthinathan, D. Kamalakkannan, R. Suresh, G. Vanangamudi, G. Thirunayraran, *Spectrochim. Acta Part A.*, 112 (2013) 245-256

[32] P. Janaki, K. G Sekar, G. Thirunayraran., *Organic Chem: An Indian J.*, 9 (2013) 68-80.

[33] R. Arulkumaran, R. Sundararajan, G. Vanangamudi, M. Subramanian, K. Ravi, V. Sathiyendiran, S. Srinivasan, G. Thirunayraran., *IUP J. Chem.*, 3 (2010) 82-98.

[34] S. Vijayakumar, R. Arulkumaran, R. Sundararajan, S. P. Sakhthinathan, R. Suresh, D. Kamalakkannan, K. Ranganathan, K. Sathiyamoorthi, V. Mala, G. Vanangamudi, G. Thirunayraran, *International Letters of Chemistry, Physics and Astronomy* 9 (2013) 68-86.

[35] G. Thirunayraran, K. G. Sekar, *International Letters of Chemistry, Physics and Astronomy* 17(2)(2014) 193-200.

[36] G. Thirunayraran, K. G. Sekar, *International Letters of Chemistry, Physics and Astronomy* 17(1) (2014) 10-20

[37] G. Thirunayraran, K.G. Sekar, R. Lakshmi Narayanan, *International Letters of Chemistry, Physics and Astronomy* 13(1) (2014) 88-94.
[38] P. Mayavel, K. Thirumurthy, S. Dinesh Kumar, G. Thirunarayanan, *Q-Science Connect*. (2014). DOI: http://dx.doi.org/10.5339/connect.2014.10

[39] G. Thirunarayanan, K. G. Sekar, *J. Taibah Univ. Sci.* 8(2014) 124-136.

[40] V. Sathiyendiran, K. G. Sekar, G. Thirunarayanan, R. Arulkumaran, R. Sundararajan, D. Kamalakkannan, R. Suresh, V. Manikanadan, R. Vijayakumar, G. Vanangamudi, *International Letters of Chemistry, Physics and Astronomy* 11(1) (2014) 33-43.

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