

# Eco-friendly Synthesis, Characterization, and Antimicrobial Evaluation of Transition Metal (II) Complexes with Thiophene-Derived Tridentate (S N N donor) Heterocyclic Schiff base Ligand

## ABSTRACT

A new unsymmetric tridentate (SNN donor) Schiff base ligand was synthesized from the 1:1M condensation of thiophene-2-carboxaldehyde with 5-amino-1,3,4-thiadiazol-2-thiol, under microwave irradiation in the presence of a green catalyst PPA-SiO<sub>2</sub>. The bivalent 3d transition metal (M = Co<sup>2+</sup>, Ni<sup>2+</sup>, & Cu<sup>2+</sup>) complexes were obtained by the reaction of M(II) chloride with tridentate Schiff base ligand (L) in a 1:2 M ratio under microwave irradiation in aqueous ethanol medium. The Schiff base and its complexes have been characterized by IR, <sup>1</sup>H NMR, UV/Vis, elemental analyses, conductometry, and magnetic measurements. The IR results showed that the Schiff base ligand acts as neutral unsymmetrical tridentate with SNN donor sequence in E isomeric form towards the metal ion through azomethine-N, thiadiazole-N, and thiophene-S. The electronic spectral results and magnetic measurement data revealed six coordinated octahedral geometry having formula [M(L)<sub>2</sub>]Cl<sub>2</sub> for the synthesized metal complexes. The melting point supported the thermal stability and non-hygroscopic nature of all the compounds. The molar conductance data revealed the 1:1 electrolytic nature of the metal complexes. The *in vitro* biological activities of the free ligand and its metal complexes against a few bacteria and fungi were screened by the disc diffusion technique. The relative order of potency of antimicrobial activity against pathogens is: Cu(II) > Co(II) > Ni(II) > Ligand.

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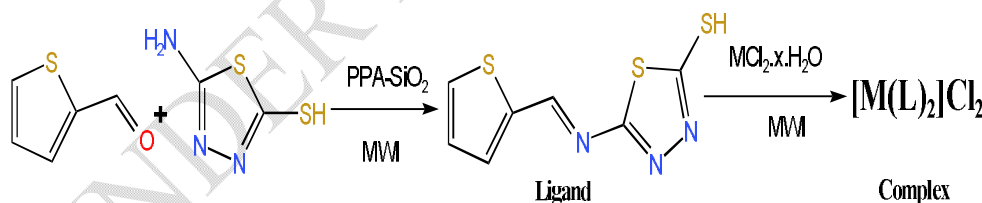
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## Graphical Abstract



**Key words:** Heterocyclic Schiff bases, Transition metal complexes, Microwave irradiation, Anti-microbial activity, Octahedral geometry

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## 1. INTRODUCTION

Schiff base (SB), also known as imine or azomethine, ligands have attracted significant interest because of their remarkable synthetic and structural characteristics, excellent tunability, chemical selectivity concerning the central metal atom/ion, unique coordination, and structural properties [1-3]. These are regarded highly promising ligands and commonly

known as 'privileged ligands' due to their remarkable ligation ability with O, N, and other donors that accommodate various geometries [4-5]. These complexes have been used as drugs and have been reported to have a wide variety of biological activities and have many biochemical, clinical, and pharmacological applications [6]. These compounds are extensively researched because they have sigma donor capabilities towards metal cations and exhibit  $\pi$  acceptor characteristics at the imine nitrogen ( $>C=N-$ ) atom, making them effective donor ligands in coordination chemistry [7]. Schiff bases, particularly those associated with heterocyclic structures, demonstrated a range of pharmacological and biological activities including antibacterial, cytotoxic, antifungal, antimalarial, anticonvulsant, antioxidant, and anti-inflammatory properties [8]. Heterocyclic Schiff base ligands with multiple donor sites, containing both nitrogen and sulfur atoms, exhibit flexible coordination capabilities with different transition metal ions [9]. Consequently, they have garnered significant attention, especially in the creation and use of bioactive coordination compounds.

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Conventional chemical synthetic methods frequently utilize solvents and chemicals that can be hazardous, posing hazards to both environmental and human health. Instead, green synthesis approaches are made to use reagents that are less hazardous to the environment and select safer solvents, or even no solvents altogether. In addition to being environmentally friendly, these methods can also lead to products that are more pure and plentiful [10-12]. Microwave-assisted synthesis can increase yields and drastically reduce reaction times from hours to only a few minutes, making it a superior alternative to conventional reflux synthesis. Microwave heating is becoming increasingly popular with chemists due to its convenience and safety features. Therefore, microwave-assisted synthesis is considered an environmentally friendly or green synthetic method, essential for modern sustainable chemistry [13-15].

The literature survey indicates that transition metal complexes with tridentate Schiff base ligands have been extensively studied and have a variety of biological applications [16]. However, little research has been done on the antimicrobial properties and green synthesis of transition metal complexes with thiophene-derived tridentate ligands [17]. The main objective of this study is to elucidate the structure and evaluate the antimicrobial properties of some newly eco-friendly synthesized Co(II), Cu(II), and Ni(II) complexes with a new tridentate (NNS donor) Schiff base which was obtained by condensation of 2-thiophenecarboxaldehyde with 5-amino-1,3,4-thiodiazol-2-thiol, under microwave irradiation in the presence of green catalyst, silica gel-supported polyphosphoric acid (PPA-SiO<sub>2</sub>) [18]. This approach expands our earlier research [19] and accounts for the various biological activities associated to new Schiff base and its complexes.

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## 2. EXPERIMENTAL STUDY

### 2.1: Materials & Methods

The starting materials 2-thiophenecarboxaldehyde and a heterocyclic amine (5-amino-1,3,4-thiodiazol-2-thiol) were obtained from Aldrich. The 3d transition metal (II) salt (CuCl<sub>2</sub>·2H<sub>2</sub>O, CoCl<sub>2</sub>·6H<sub>2</sub>O & NiCl<sub>2</sub>·6H<sub>2</sub>O) and other chemicals used in this research project were obtained from Aldrich and stored in controllable conditions in a glovebox. Solvents like ethanol (EtOH), N, N-dimethylformamide (DMF), dimethyl sulfoxide (DMSO), acetone, and molecular sieves (0.4 nm) were purchased from Merck.

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A modified microwave oven model 2001 ETB (Bajaj Electricals Limited, Mumbai, India) with rotating tray and a power source 230V, microwave energy put out 800W and microwave frequency 2450 MHz was used for green synthesis of ligand as well as undertaken M(II) complexes. The progress of the synthetic reaction was monitored by performing TLC using Al sheets precoated with Merck 60 F<sub>254</sub> that was visualised using a UV lamp.

With EA-1106 elemental analyzer (Carlo-Erba) the elemental analysis of C, H, and N content and empirical formulas of the compounds / complexes were determined. The metal contents were estimated using standard methods. The chloride content in the complexes was determined gravimetrically as silver nitrate test.

Molar conductance ( $10^{-3}$  M) was measured by Elico-conductivity bridge at room temperature. The magnetic susceptibility measurements were carried out on a Gouy balance at room temperature using Hg[Co(SCN)<sub>4</sub>] as the calibrant. Electronic spectra were recorded (in DMSO at  $10^{-3}$  M) on a Shimadzu UV-Vis-160A spectrophotometer. Shimadzu 8400-S FT-IR spectrophotometer ( $\lambda = 4000-400$  cm<sup>-1</sup>) was used to record the infrared spectra using KBr pellets. Bruker Avance Digital 500-NMR spectrometer (operating at 500 MHz) was used for recording the <sup>1</sup>H NMR spectra in DMSO-d<sub>6</sub> using TMS as an internal standard. The chemical shift was measured in ppm on the  $\delta$  scale and the coupling constants were measured in Hertz.

## 2.2: Preparation of Green Catalyst (PPA-SiO<sub>2</sub>)

SiO<sub>2</sub> (5g 200-400 mesh) was added to the polyphosphoric acid (PPA) solution in chloroform at 320K and stirred for one hour until solid formation. To remove the unused chloroform, a rotary evaporator was used and the resulting solid was dried under vacuum at 298K [20].

## 2.3: Green Synthesis of Heterocyclic Schiff Base Ligands

A mixture in equimolar proportions of 2-thiophenecarboxaldehyde (I) and a 0.01 molar ethanolic solution of heterocyclic amine [5-amino-1,3,4-thiadiazole-2-thiol (II)] along with PPA-SiO<sub>2</sub> in each scenario was exposed to microwave irradiation for 10 minutes at 360W and 360K, with the reaction process being tracked using TLC. (scheme-1). The reaction mixture was maintained at room temperature and then added cooled water while stirring for a few minutes, resulting in the production of a bright yellow powder solid product that was collected by suction filtration. The pure form of the ligand was obtained by recrystallization with aqueous ethanol and identified or characterized by spectral studies and the melting point determination [yield = 95% (conventional 82%)].

## 2.4: Green Synthesis of Transition metal (II) Complexes

An aqueous-ethanolic solution of the ligand (L) and the respective transition metal (II) chloride was thoroughly mixed in 1:2 (metal:ligand) stoichiometric ratio and irradiated in the microwave oven adding 0.1% ethanolic KOH solution to maintain the pH within the range of 7-8 for a period of 5-7 minutes, which produced the better yield (80-85%) of the respective metal (II) complexes (scheme-1) compared to the conventional method (55-60%). Thus, obtained coloured solid products were filtered and recrystallized from DMF, washed with ethyl acetate, and then dried under reduced pressure over anhydrous CaCl<sub>2</sub> in a desiccator.

## 2.5: Antimicrobial Activity of Ligands and Metal (II) Complexes

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The *in vitro* antimicrobial (anti-bacterial and anti-fungal) activities of the investigated ligand and its transition metal (II) complexes were evaluated by disc-diffusion method [21-23]. The *in vitro* anti-bacterial activity of the undertaken compounds was achieved against two Gram-positive bacteria {*Staphylococcus aureus* (SA) & *Bacillus subtilis* (BS)} and two Gram-negative bacteria {*Escherichia coli* (EC) & *Salmonella typhi* (ST)} using chloramphenicol as standard reference. The anti-fungal activity of the undertaken compounds was performed against fungal strains {*Candida albicans* (CA) & *Aspergillus niger* (AN)} using griseofulvin as standard reference of the same concentration (20 µg/mL) under identical conditions.

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Accurately weighed 10 mg each of the green synthesized ligand and its transition metal (II) complexes was dissolved separately in DMSO and volume made up to 10 mL in volumetric flask (1.0 mg/mL). These solutions were further diluted with DMSO to appropriate concentration (100 µg/mL). Similarly, solutions of 20 µg/mL of concentration of each chloramphenicol and griseofulvin were prepared from their stock solutions. Nutrient broth, for antibacterial activity, has composition of 2.0 g yeast extract, 1.0 g beef extract, 5.0 g peptone and 5.0 g of NaCl in distilled water and volume made up to 1L. Nutrient Agar medium was obtained by adding 2.0% of agar to nutrient broth at 7.4 pH. For antifungal activity, Sabouraud dextrose medium has composition 10.0 g peptone, 40.0 g dextrose in 1.0 L distilled water at 5.7 pH and agar medium was obtained by adding 1.5% of agar to it.

The sterile disks were drenched in investigated test compounds and were carefully placed on incubated agar surface separately. The petri-dishes were incubated for 24 hours at 37 K for bacteria and for 48 hours at 37 K for fungal strain. Finally, the zone of inhibition (in mm) was carefully measured separately for each compound tested by comparing them with standard drugs. Each test was performed in triplicate in an individual experiment, and the mean is reported (table-4). The various concentrations of test compounds in DMSO like 50, 30, and 20 µg/mL were first prepared and tested for inhibition. Further dilutions were made and tested till the concentration inhibition was observed. The broth dilution method was used to determine the minimum inhibitory concentration (MIC) against the mentioned bacteria [24]. The observations related to MIC are presented in table-5.

### 3. RESULTS & DISCUSSION

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#### 3.1: Synthesis & Characterization

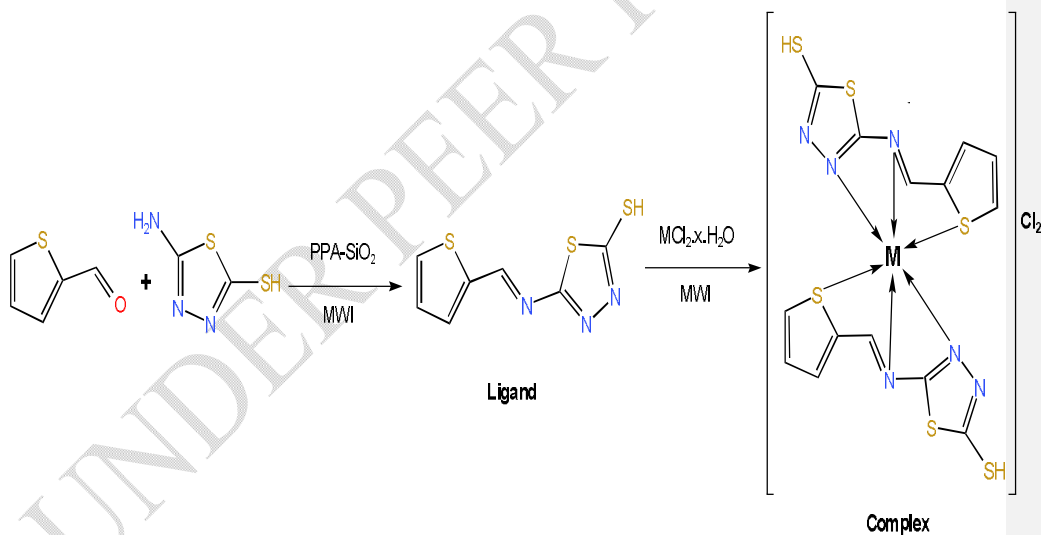
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The heterocyclic Schiff base (L) was synthesized using a microwave-assisted green approach by condensation of heterocyclic aldehyde (2-thiophenecarboxaldehyde, I) with thiadiazole amine [(II)] in the presence of PPA-SiO<sub>2</sub> as a solid acid catalyst under clean and environmentally benign conditions. The condensation involves the nucleophilic attack of NH<sub>2</sub> group on the electrophilic H-C=O group followed by dehydration to form corresponding Schiff bases. The used catalyst (PPA-SiO<sub>2</sub>) was recycled by simple filtration and can be used repeatedly; therefore, it acts as a green catalyst and the present synthetic approach complies with the principle of sustainable chemistry [25]. The M(II) complexes were formed by direct ligation of ligand (L) in the appropriate molar to metal precursors ratio 1:2 (metal:ligand) and

furnished the corresponding complexes in better yields (80-85% from 57-60%). This is probably due to the increased homogeneity of reaction mixture by rotation of the reaction platform tray in the microwave oven. Obviously, the microwave-assisted synthesis of mentioned ligand and its respective M (II) complexes were completed in shorter time with higher yields compared to the conventional synthetic methods. The synthetic reaction completion time and the yields of the products using a green synthetic approach and conventional method are presented in **table -1** for comparison.

### 3.2: Physico-analytical Studies

The heteronuclear Schiff base and all the eco-friendly synthesized transition metal (II) complexes are stable and non-hygroscopic coloured solid at room temperature. The complexes have range of melting points or decomposition temperatures. The complexes are insoluble in common organic solvents but soluble in DMF and DMSO. The elemental analysis of the synthesized M(II) complexes  $\{[ML_2]Cl_2\}$  also verifies % composition of C, H, and N constituents, and agreed with the proposed structures corresponding to  $\{1:2$  (metal: ligand) $\}$  stoichiometry and mononuclear nature. The values of observed molar conductance of the metal (II) complexes ( $88.54$  to  $98.60 \text{ ohm}^{-1}\text{cm}^2\text{mol}^{-1}$ ) were appreciable and to account for their any dissociation in DMF at room temperature, which revealed their 1:1 electrolytic nature [26].



**Scheme-1:** Microwave-assisted green synthesis of ligand and its metal(II) complexes

**Table-1:** Microanalytical & physical data with the comparative results of conventional and green synthetic methods of the investigated compounds

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Molecular formula of compounds (colour)	Molecular wt. (g mol <sup>-1</sup> )	Melting Pt. (K)	Elemental analysis % calculated (found)				Molar Conductance (ohm <sup>-1</sup> cm <sup>2</sup> mol <sup>-1</sup> )	Reaction Time CM (GM)	% Yield CM (GM)	
			C	H	N	M				
C <sub>7</sub> H <sub>5</sub> N <sub>3</sub> S <sub>3</sub> Bright yellow	227.3	458	37.0 (37.2)	2.2 (2.5)	18.5 (18.6)	-	-	2 h (10 min.)	80	Comment [DS223]: range?
[CoL <sub>2</sub> ]Cl <sub>2</sub> Light pink	584.0	490	28.8 (28.5)	1.7 (1.5)	14.4 (14.5)	10.09 (10.0)	93	1h (5 min.)	58	Comment [DS224]: range
[NiL <sub>2</sub> ]Cl <sub>2</sub> Light green	583.7	494	28.8 (28.5)	1.7 (1.9)	14.4 (14.5)	10.05 (10.02)	96	1h (6 min.)	56	Comment [DS225]: range
[CuL <sub>2</sub> ]Cl <sub>2</sub> Green	588.5	486	28.5 (28.4)	1.7 (1.5)	14.3 (14.5)	10.79 (10.80)	90	1h (7 min.)	55	Comment [DS226]: range

CM = Conventional method, time in hours; GM = Green method, time in minutes

### 3.3: Spectroscopic Characterization

#### 3.3.1: Infrared Spectral Studies

The infrared spectral data of the investigated compounds {ligand & metal(II) complexes} are presented in Table 2. Comparison of the IR spectra of the ligand and that of metal (II) complexes reveals the involvement of coordination sites in chelation. The IR spectral data of the metal (II) complexes contained all the major absorption bands of the ligand and some new bands indicative of the tridentate coordination of the ligand with the metal (II) ions in the complexes.

The IR spectrum of the undertaken Schiff base (L) exhibited the azomethine (HC=N) and thiadiazole stretching at 1640 and 1620 cm<sup>-1</sup> respectively. The thiol (S-H) stretching appeared in the 2590 cm<sup>-1</sup> region along with a band resulting from the thiophene ring (ν<sub>C-S-C</sub>) stretching of thiophene moiety at 850 cm<sup>-1</sup> [27].

Comparison of the infrared spectra of the Schiff base ligand was performed as a reference and their metal (II) complexes showed that the ligand was principally tridentate coordinated with the central metal (II) ions. The absorption band appearing at 1640 cm<sup>-1</sup> due to azomethine stretching vibration is shifted to a lower frequency, indicating the participation of azomethine nitrogen in complexation [28]. The absorption band at 1620 cm<sup>-1</sup> assigned to the thiadiazole ring (C=N) vibration is also shifted to lower frequency, which is indicative of the involvement of the thiadiazole ring in complexation [29]. Furthermore, an absorption band at 850 cm<sup>-1</sup> attributed to thiophene ring C-S-C in the ligand shifted to lower frequency in the respective metal (II) complexes. This shows the sulfur in the thiophene is coordinated in the complexation. Finally, conclusive evidence of coordination of the ligand with metal atoms was established by far IR spectra in which new absorption bands at 525, 530, and 540 cm<sup>-1</sup> assigned to M-N (azomethine), M-N (thiadiazole), and M-S (thiophene S) were observed in the spectra of the metal (II) complexes, which were not present in the infrared spectra of the ligand.

Comment [DS227]: Fourier Transform Infrared Spectroscopy (FTIR)

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**Table-2:** Observed IR bands ( $\text{cm}^{-1}$ ) of Schiff bases & its metal (II) complexes

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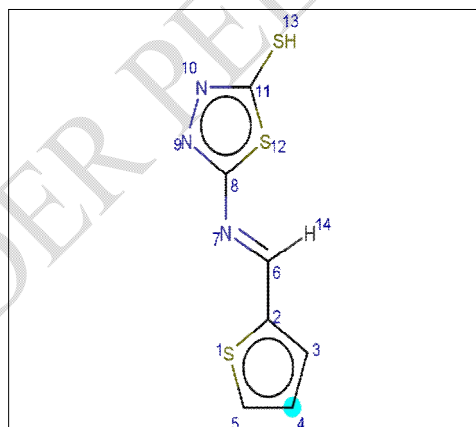
Compound	$\nu(\text{HC}=\text{N})_{\text{azomethine}}$	$\nu(\text{C}=\text{N})_{\text{thiadiazole}}$	$\nu(\text{S}-\text{H})_{\text{thiophene}}$	$\nu(\text{N}-\text{M})_{\text{azomethine}}$	$\nu(\text{N}-\text{M})_{\text{thiadiazole}}$	$\nu(\text{S}-\text{M})_{\text{thiophene}}$
(L)	1635	1620	850	-	-	-
$[\text{Co}(\text{L})_2]\text{Cl}_2$	1625	1610	835	525	525	360
$[\text{Ni}(\text{L})_2]\text{Cl}_2$	1625	1610	835	530	530	365
$[\text{Cu}(\text{L})_2]\text{Cl}_2$	1620	1605	835	525	525	365

Obviously, the infrared spectral studies of the ligand and its M(II) complexes revealed that the investigated Schiff base ligand acted as a neutral tridentate (designated as SNN donor) ligand.

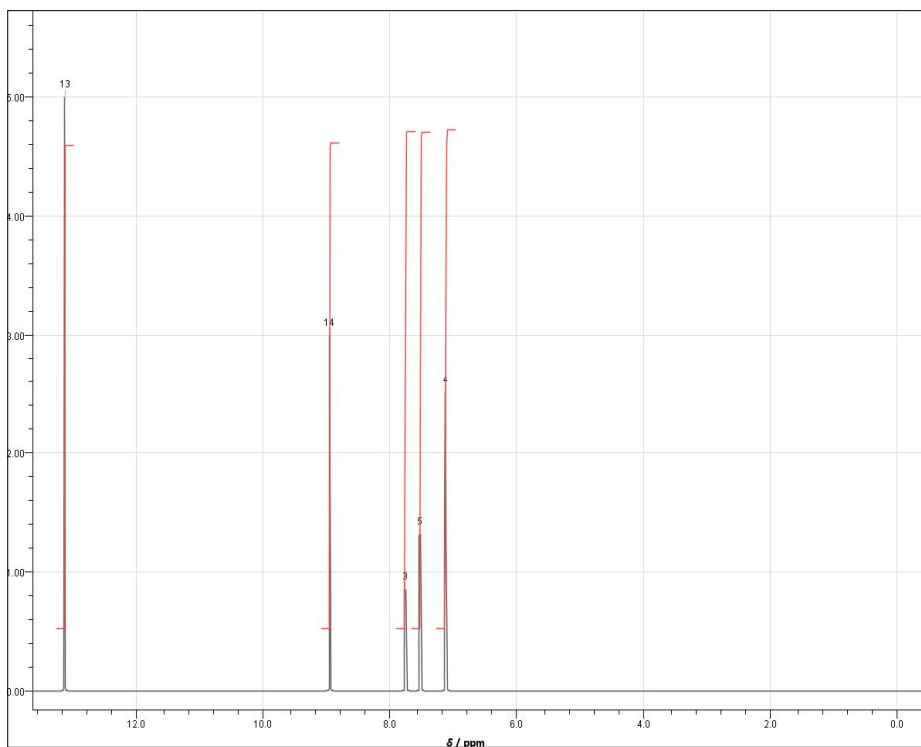
### 3.3.2: $^1\text{H}$ NMR Spectral Studies

The  $^1\text{H}$  NMR spectra of the investigated Schiff base displayed the SH proton as singlet at  $\delta$  13.12 ppm, the azomethine ( $\text{HC}=\text{N}$ ) proton appeared as singlet at  $\delta$  9.05 ppm.  $\text{C}_5\text{-H}$  and  $\text{C}_3\text{-H}$  of the thiophene moiety appeared as doublets at  $\delta$  7.59 ppm and  $\delta$  7.82 ppm respectively. Similarly,  $\text{C}_4\text{-H}$  appeared as a double doublet at  $\delta$  7.127 ppm. (Figure-1&2).

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**Figure-1:** Structure of the ligand



**Figure-2:**  $^1\text{H}$  NMR spectrum of the investigated Schiff base (L)

### 3.3.3: Magnetic Property & Electronic Spectral Studies

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Electronic absorption (UV-vis) spectral studies along with the observed magnetic moment values of the metal (II) complexes are quite useful for the nature of the ligand field around the metal ion and thus can be used for the prediction of probable geometry or stereochemistry in association with other spectroscopic methods. It is well established fact that the extension of the absorption spectra from UV region to visible region is due to ligand to metal charge transfer (LMCT) and d-d transition bands of the metal in the complexes. The structure of the coordination complexes can be assigned based on the position and number of peaks of the *d-d* transitions in their electronic spectra. The observed *d-d* transitions and determined magnetic moment values of the investigated metal (II) complexes with their plausible geometry are presented in table-3.

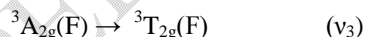
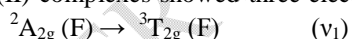
The ligands show two bands at absorption bands at 252 and 310 nm assigned for  $\pi \rightarrow \pi^*$  and  $n \rightarrow \pi^*$  electronic transitions respectively.

**Table-3:** Observed electronic bands, magnetic moments, and geometry of M(II)-complexes

M-complexes	d-d transition bands (nm)	$\mu_{\text{eff}}$ (BM)	Stereochemistry of M(II) Complexes
[Co(L) <sub>2</sub> ]Cl <sub>2</sub>	328, 548, 1141	4.4	Octahedral
[Ni(L) <sub>2</sub> ]Cl <sub>2</sub>	352, 616, 987	3.4	Octahedral
[Cu(L) <sub>2</sub> ]Cl <sub>2</sub>	326, 448	1.4	Distorted octahedral

The electronic spectra of the investigated Co(II) complexes show three absorption bands corresponding to the  ${}^4T_{1g} \rightarrow {}^4T_{2g}$  (F),  ${}^4T_{1g} \rightarrow {}^4T_{2g}$  (P) and  ${}^4T_{1g} \rightarrow {}^3A_{2g}$  (F) electronic transitions respectively suggesting the octahedral geometry which is substantiated by the observed magnetic moment values (4.3-4.4 BM) corresponding to three unpaired electrons per Co(II) ion in an octahedral environment [30].

The electronic spectra of Ni(II) complexes showed three electronic absorption bands assigned for



electronic transitions which suggested the octahedral geometry of the metal complexes. The determined magnetic moment values of the Ni (II) complexes were 3.3-3.4 BM which correspond to two unpaired electrons per Ni<sup>2+</sup> ion for the ideal six-coordinated octahedral configuration [31].

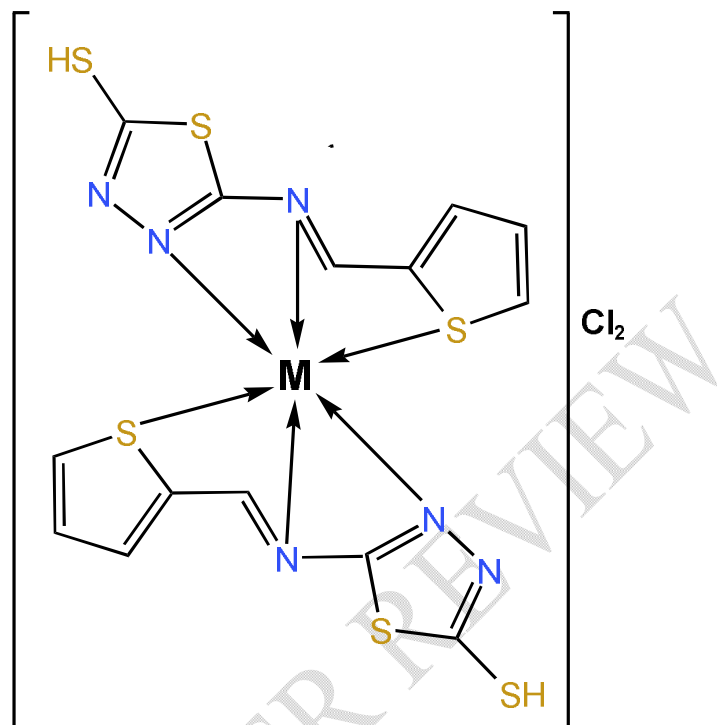
The electronic spectrum of the investigated Cu(II) complex showed broad absorption bands between 10Dq band for octahedral geometry corresponding to the  ${}^2E_g \rightarrow {}^2T_{2g}$  transition. The determined magnetic moment values of the Cu(II) complexes were 1.3-1.4 BM, which are indicative of one unpaired electron per Cu<sup>2+</sup> ion suggesting that the investigated complexes had structure within the range consistent with spin-free distorted octahedral geometry [32].

### 3.4: Proposed Structure of the M(II)-Complexes

Based on the aforesaid experimental evidences, we tentatively propose the octahedral /distorted octahedral structure of the investigated M(II)-complexes (where M = Co, Ni, & Cu) in which the heterocyclic Schiff base ligands act as a neutral tridentate (NNS Donor) ligand.

**Figure-3** presents the common tentative octahedral geometry/structure of the complexes.

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**Figure-3:** Proposed octahedral structure of the investigated M(II) complexes

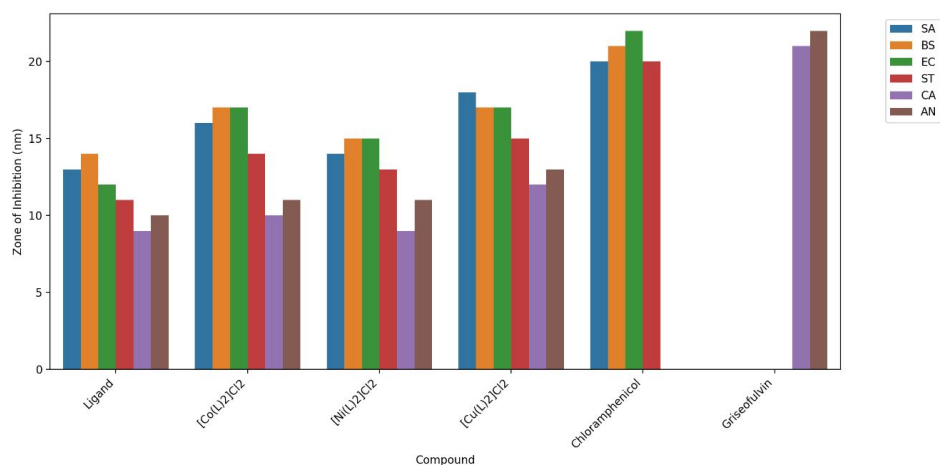
### 3.5: Antimicrobial Activities

The experimental results of the antimicrobial activity of the undertaken heterocyclic Schiff base ligands and its transition metal (II) complexes are presented in table-4 and table-5.

**Table-4:** Determined zone of inhibition (in nm) of investigated compounds against pathogens

Compound	Concentration	Zone of inhibition in nm					
		Gram-(+)-bacteria		Gram(-)-bacteria		Fungi	
		SA	BS	EC	ST	CA	AN
Ligand	100 µg/mL	13	14	12	11	9	10
[Co(L) <sub>2</sub> ]Cl <sub>2</sub>	100 µg/mL	16	17	17	14	10	11
[Ni(L) <sub>2</sub> ]Cl <sub>2</sub>	100 µg/mL	14	15	15	13	9	11
[Cu(L) <sub>2</sub> ]Cl <sub>2</sub>	100 µg/mL	18	17	17	15	12	13
Chloramphenicol	20 µg/mL	20	21	22	20	-	-
Griseofulvin	20 µg/mL	-	-	-	-	21	22
DMSO	-	-	-	-	-	-	-

Figure 4 : Zone of inhibition

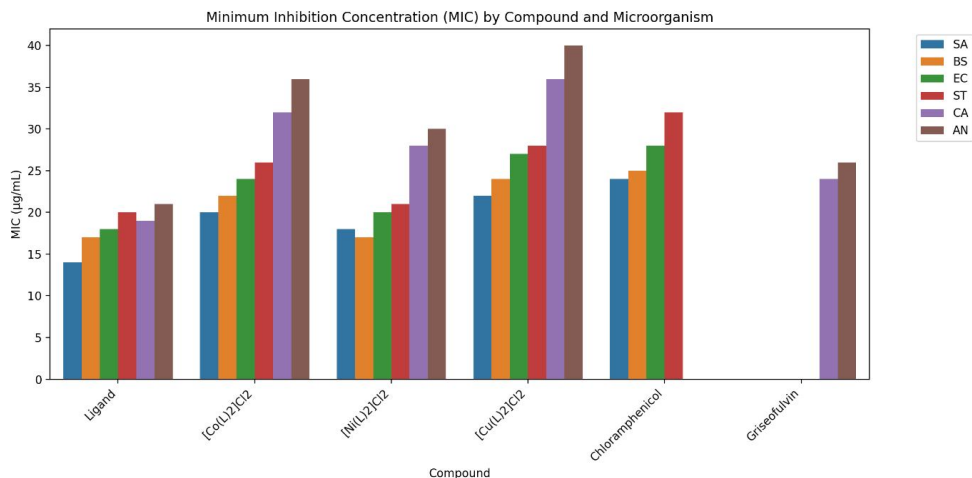


**Table-5:** Minimum Inhibitory Concentration (MIC) of compounds against pathogens

Compound	Minimum inhibition Concentration (MIC) in $\mu\text{g/mL}$					
	Gram-(+)-bacteria		Gram-(-)-bacteria		Fungi	
	SA	BS	EC	ST	CA	AN
Ligand	14	17	18	20	19	21
[Co(L) <sub>2</sub> ]Cl <sub>2</sub>	20	22	24	26	32	36
[Ni(L) <sub>2</sub> ]Cl <sub>2</sub>	18	17	20	21	28	30
[Cu(L) <sub>2</sub> ]Cl <sub>2</sub>	22	24	27	28	36	40
Chloramphenicol	24	25	28	32	-	-
Griseofulvin	-	-	-	-	24	26

Figure .5

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The data and bar plots of antimicrobial activities of the investigated compounds showed that the metal complexes have enhanced activity compared to the free ligand against the same micro-organisms under the identical experimental conditions. The metal complexes show moderate activity compared to standard drugs despite being at higher concentrations [33]. [Cu(L)<sub>2</sub>]Cl<sub>2</sub> shows the best overall antimicrobial activity among the metal complexes. The order of activity for metal complexes is :



Furthermore, the gram-positive bacteria (SA, BS) show slightly higher susceptibility than gram-negative bacteria (EC, AN) and fungi (CA, AN) generally show lower zones of inhibition compared to bacteria.

The metal complexes maintain activity across all tested organisms, showing broad-spectrum antimicrobial properties. The enhancement in antimicrobial activity of metal complexes compared to the free ligand suggests successful coordination and improved bioactivity through complexation[34]. Apart from this, other factors such as solubility, conductivity, and dipole moment (influenced by the presence of metal ion) may also be the possible reasons for enhancing their antimicrobial activity [35].

## CONCLUSIONS

An efficient and facile synthesis of a new pharmaco-active ligand in a single step by PPA-SiO<sub>2</sub>-catalysed condensation of 2-thiophenecarboxaldehyde with 5-amino-1,3,4-thiodiazol-2-thiol in an ethanolic solution as a green solvent and its bivalent transition metal complexes bearing possible pharmacophore were obtained under microwave irradiation with higher yields in less time compared to conventional heating. The ligand coordinated to the M(II) ion in a neutral tridentate (SNN donor) mode. The octahedral geometry of the metal complexes{[M(L)<sub>2</sub>]Cl<sub>2</sub>} has been proposed based on elemental analyses, molar conductance,

magnetic measurement, and electronic spectral studies. The antimicrobial activity studies indicated that the M (II) complexes possess comparatively higher activities than the free ligand owing to enhanced lipophilicity by chelation. The order of antimicrobial activity for compounds is :

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Cu-complex > Co-complex > Ni-complex > Ligand

The new synthesized compounds were found to be more active against Gram-positive than Gram-negative bacteria. The results of these analyses indicated the substantial potential of green synthesized metal complexes in the biological field as future drugs.

The present eco-friendly synthetic procedure represents an alternative to the existing methods for the synthesis of bioactive thiophene-derived Schiff base ligand and its metal (II) complexes under microwave irradiation. This article describes a simple, proficient, and sustainable approach to synthesis with easy workup, higher yield, and economic viability compared to other methodologies.

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Authors hereby declare that no generative AI technologies such as Large Language Models (ChatGPT, COPILOT, etc) and text-to-image generators have been used during writing or editing of manuscripts.

#### DATA AVAILABILITY STATEMENT

The data that have been used are present here.

#### ETHICAL STATEMENT

The authors declare that the research for the manuscript submitted does not involve human or animal research subjects.

#### REFERENCES

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