



## INVITRO ANTIBACTERIAL AND ANTIFUNGAL ACTIVITIES OF NITRITO-O COMPLEX OF Hg(II) WITH BENZIMIDAZOLE

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### ABSTRACT

Transition metal complex of Benzimidazole (Bz) and nitrito-O (NO<sub>2</sub>) ligands was synthesized and characterized based on elemental analysis, metal estimation, molar conductance, cyclic voltammetry, solid state reflectance (UV-Spectra), IR, Far-IR and NMR spectra. The data obtained from elemental analysis, metal estimation and molar conductance the proposed general formula of the complex is [M(L<sub>1</sub>)<sub>2</sub>(L<sub>2</sub>)<sub>2</sub>] where M = Hg(II), L<sub>1</sub> – Benzimidazole and L<sub>2</sub> – Nitrite ion ( Nitrito-O). The complex is neutral in nature. Solid state reflectance depicts the tetrahedral geometry of the complex. Metal- chelate and complexation ability confirmed by their IR and Far-IR spectral data. Nuclear magnetic resonance spectra (<sup>1</sup>H-NMR & <sup>13</sup>C-NMR) conveniently predict the geometry of the complex. The mixed ligand complexes were evaluated for their biological activities against the bacterial strain *Klebsiella Pneumonia* and the fungal strain *C.albicans*. The complex was found to very good antibacterial and antifungal activities compared to the Benzimidazole ligand.

**KEYWORDS:** Hg(II) complex, Benzimidazole, Nitrito-O, *Klebsiella Pneumonia* and *C.albicans*.

### INTRODUCTION

Nitrogen and oxygen based ligands are extensively used for the coordination complexes of transition metal ions. Benzimidazole is one the heterocycles which exhibit diverse biological activities.<sup>[1]</sup> benzimidazole derivatives were used to antibacterial, antifungal, antiviral, antioxidant, anthelmintic, anti-inflammatory, anticancer, antihistaminic, anti-anxiety and anti hypertensive activities.<sup>[2-4]</sup> microwave assisted organic synthesis is simple, clean, fast efficient and economic.<sup>[5-6]</sup> microwave assisted synthesis have some advantages in the field of multistep total synthesis, medicinal chemistry, drug discovery, material science and nano technology.<sup>[7-9]</sup> in this paper we have reported microwave assisted synthesis, spectral and biological activities of Hg(II) complex with Benzimidazole and nitrite ion ligands.

### MATERIAL AND METHODS

All the chemicals used were of AR grade from Alfa Aesar and were used as such. Microwave irradiations were used for the synthesis of complexes from domestic microwave oven. The elemental analysis of the complex was carried out using (Thermo Finnegan make, Flash EA1112 series) CHNS(O) analyzer instrument. The metal ion estimated by gravimetrically using standard procedure. The molar conductance of 10<sup>-3</sup> M complex in acetonitrile was conducted using Systronic Conductivity Bridge at 25<sup>o</sup>C. The cyclic voltammetry of the complex

were carried out using Princeton applied research cyclic voltammogram. The diffused reflectance spectra of the complex in the solid state were measured using Varian carry-5000 model UV-Visible spectrophotometer. IR spectra of the free Benzimidazole and its complex were carried out using Shimadzu FT-IR8400s spectroscopy at 4000-400 cm<sup>-1</sup> wave number with KBr pellet technique. The Far IR spectrum of the complex was recorded in a Bruker make, 3000 Hyperion Microscope with Vertex 80 FTIR system model instruments.

The antibacterial and antifungal activities of Benzimidazole and its complex were done by in-vitro Agar well diffusion method using Amikacin and ketoconazole as a standard for bacterial and fungal strain respectively.

### Preparation of complexes

The Hg(II) complex was synthesized by mixing Benzimidazole 0.87 g (7.36 mmol) in 10ml methanol to the mercury chloride 1g (3.68 mmol) in 10ml methanol. The mixture was irradiated on a microwave oven for 10 sec, then sodium nitrite 0.50g (7.36 mmol), in 10 ml ethanol was added to the above mixture respectively. The whole mixture was irradiated on a microwave oven for another 10 sec. The precipitated colourless complex was filtered and washed with ethanol, dried in vacuum desiccators and kept in an airtight glass contained.

## RESULTS AND DISCUSSION

### i) Micro-analytical data

The elemental analysis and metal estimation of the prepared complex indicate their molecular formula. The metal and ligands are present in the complex by the ratio of (1:2:2) metal: Benzimidazole: nitrite is the

predominant evidence for the formula of the complex. The complex have the formula of  $[\text{Hg}(\text{Bz})_2(\text{NO}_2)_2]$  which is monomeric in nature. The molar conductance value is  $21.80 \text{ Ohm}^{-1}\text{cm}^2\text{mol}^{-1}$  in acetonitrile were measured, according to Geary *et al.*, the complex exhibit non-electrolyte (1:0 type) neutral<sup>[10]</sup> complex.

**Table 1: Micro analytical data.**

S.No.	Complex	Colour	% Yield	Molecular weight (g/mol)	Elemental analysis				
					%C	%H	%N	%O	%M
1	$[\text{Hg}(\text{Bz})_2(\text{NO}_2)_2]$	colourless	65	574.85	29.22	2.08	14.61	11.13	34.89

### ii) Solid state reflectance spectra

The solid state diffused reflectance spectra of the Hg(II) complex gives only charge transfer spectra at  $36764 \text{ cm}^{-1}$  (272 nm). 5d orbital of metal ion is completely filled; there are no unpaired electrons to taking electronic transitions and their fore C-T band only is observed.<sup>[11]</sup>

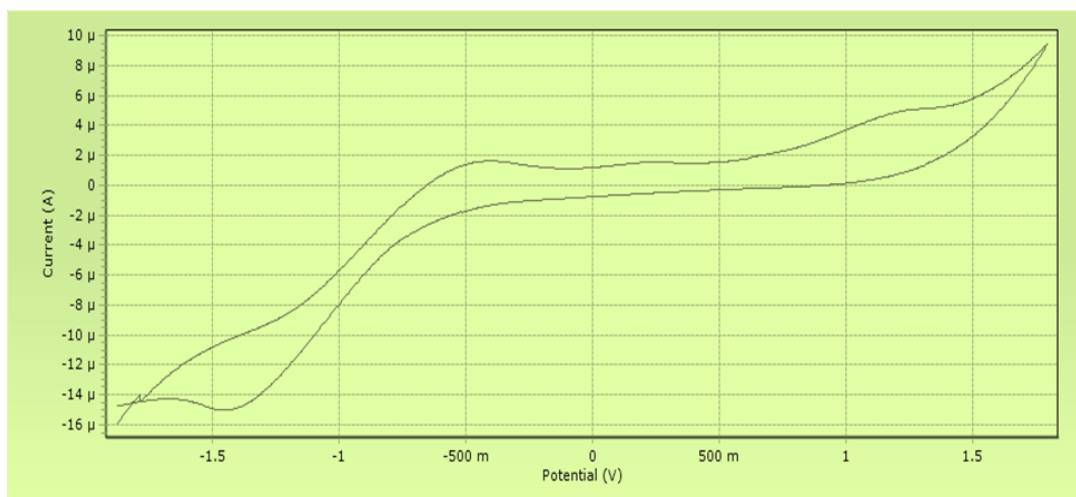
### iii) IR and Far-IR Spectra

Complexing ability and metal chelation of the complex predicted using IR and Far-IR spectral data. IR spectra of free Benzimidazole exhibits strong stretching frequencies at  $1400\text{-}1650 \text{ cm}^{-1}$  indicating the  $\nu(\text{C}=\text{N})$  of Benzimidazole ring which is shifted to  $1672 \text{ cm}^{-1}$  in the complex may exhibit the metal ion can coordinated through the nitrogen atom.<sup>[12]</sup> The other frequencies  $\nu(\text{N}-\text{H})$  at  $3200\text{-}3100 \text{ cm}^{-1}$ ;  $\nu(\text{C}-\text{N})$  at  $1270\text{-}1290 \text{ cm}^{-1}$  and  $\nu(\text{C}-\text{H})$  aromatic, at  $3100\text{-}3300 \text{ cm}^{-1}$ ; are shifted to  $3305 \text{ cm}^{-1}$ ,  $1275 \text{ cm}^{-1}$  and  $3080 \text{ cm}^{-1}$  respectively indicating

the Benzimidazole can enter into the coordination sphere. The mixed ligand nitrite ion shows  $\nu_a(\text{NO}_2)$  at  $1234 \text{ cm}^{-1}$  and  $\nu_s(\text{NO}_2)$  at  $1362 \text{ cm}^{-1}$  were also confirming the additional ligand also enter into the sphere.<sup>[13]</sup> metal chelating ability of complex also predicted and confirming the  $\nu(\text{M}-\text{N})$  and  $\nu(\text{M}-\text{O})$  coordination.<sup>[14]</sup> Hg-N (Imidazole nitrogen atom of C=N) at  $435 \text{ cm}^{-1}$  and Hg-O (Nitrito-O of  $\text{NO}_2$  ion) at  $346 \text{ cm}^{-1}$  confirming metal chelation from Far-IR spectra.

### iv) Cyclic voltammeter

Mercury metal ions have different oxidation state (+2, +1 0). The metal ion reduction and oxidation properties was perfectly depicted from cyclic voltammeter study.<sup>15</sup> the cyclic voltammogram of Hg(II) complex show one electron quasi reversible reaction of Hg(II)/Hg(I) couple with Epc at  $-1.235\text{V}$  and Epa at  $-0.500\text{V}$  and  $\Delta E_p = -0.735\text{V}$ .<sup>[15]</sup>



**Figure-2: Cyclic voltammogram of Hg(II) complex.**

### v) $^1\text{H}$ and $^{13}\text{C}$ NMR spectra

Proton NMR and carbon NMR spectra of the diamagnetic Hg(II) complex were carried out in DMSO- $d_6$  using Tetramethylsilane as the internal standard. The  $^1\text{H}$ -NMR spectra of free Benzimidazole shows chemical shift values at 12.53 ppm, 8.25 ppm and 7.17-7.20 ppm corresponding to N-H, H-C=N and aromatic C-H groups respectively these are shifted to downfield or up field at 12.58 ppm, 8.42 ppm and 7.19-7.20 ppm respectively in the complex confirming the complexation nature of

ligands to the metal ions.<sup>[16]</sup> in  $^{13}\text{C}$ -NMR of Benzimidazole gives the chemical shift values at 142.42 ppm, 122.20 ppm, 118.20 ppm and 113.07 ppm corresponding to four different carbon of ligand which are downfield/up field in the complex at 150.43 ppm, 140.10 ppm, 122.73 ppm and 116.88 ppm respectively.<sup>[17]</sup>

### vi) Bio-potential activities

#### Antibacterial and antifungal activities

The Invitro antibacterial and antifungal activities of Hg(II) complex were done by Agar well diffusion method using Amikacin and ketoconazole as the standards and DMSO as the solvent control. Metal-

chelates exhibit significant activities against *Klebsiella Pneumonia* and the fungal strain *C.albicans* than the ligand Benzimidazole. However chelate nature and neutral nature of complex shows its higher activity than the ligand.



Figure-3: Inhibition of Hg(II) and Benzimidazole of *Klebsiella Pneumonia* and *C.albicans*

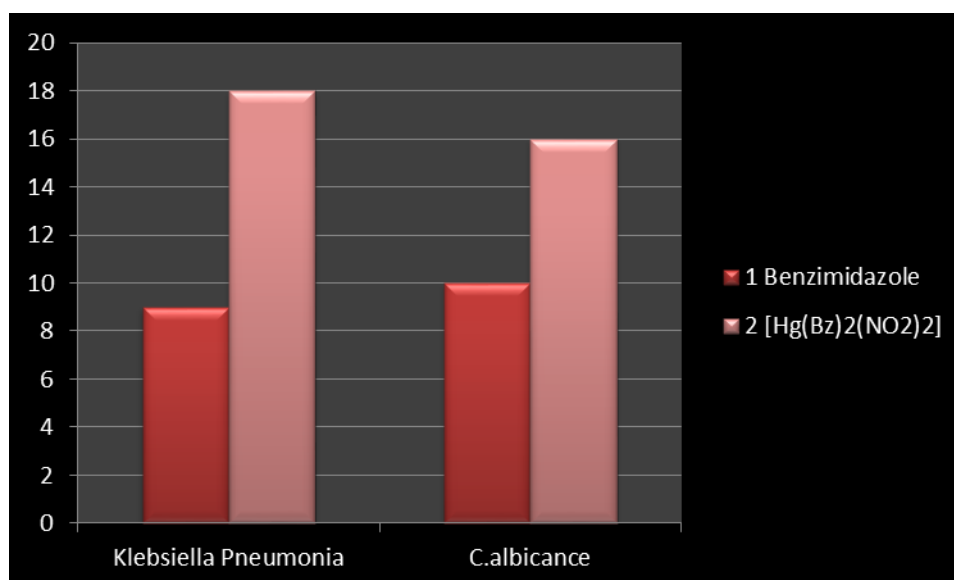


Figure 4: Comparative Bio-potential activities of Hg(II) and Benzimidazole of *Klebsiella Pneumonia* and *C.albicans*.

### CONCLUSION

In this effort, the Hg(II) complex of Benzimidazole and nitrite ion was synthesized using microwave irradiated method and characterized on the basis of analytical, spectral, and Biological methods. The formulae of the Hg(II) complex  $[\text{Cu}(\text{Bz})_2(\text{NO}_2)_2]$  and they are non-ionic, neutral and non-electrolyte. The solid state diffused reflectance spectra reveal that the complex has tetrahedral geometry. The bio-potential activities show the complex is biologically active against the tested microorganisms than the free ligand.

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### REFERENCES

1. Synthesis of novel dipodal-benzimidazole, benzoxazole and benzothiazole from cyanuric Chloride: Structural, photophysical and antimicrobial studies, Vikas S. Padalkar, Vinod D. Gupta, Kiran R. Phatangare, Vikas S. Patil, Prashant G. Umape, N. Sekar, Journal of Saudi Chemical Society, 2014; 18: 262-268.
2. Catalytic assemble of 2-amino-5,6-dimethyl-1H-benzimidazole with carboxylic acids and evaluation of their antimicrobial and antioxidant activities, Aruna Sindhe M., Yadav D. Bodke, Kenchappa R.,

- Sandeep Telkar, Chandrashekar A., and Vinoda B.M., Journal of Taibah University Medical Sciences, 2016; 11(5): 418-426.
3. synthesis, molecular docking and biological evaluation of imides, pyridazines, and imidazoles derived from itaconic anhydride for potential antioxidant and antimicrobial activities, Prakash S. Nayak, B. Narayana, B. K. Sarojini, Sana Sheik, K. S. Shashidhar, K. R. Chandrasheka, Journal of Taibah University for Science, 2016; 10: 823–838.
  4. Benzimidazole: as potential biologically active agent, Vikash Kumar Chaudhari, Devender Pathak and Satyendra Sing nt. Res. J. Pharm, 2014; 5(12): 861-875.
  5. Synthesis and antimicrobial activity of novel 2-substituted benzimidazole, benzoxazole and benzothiazole derivatives, Vikas S. Padalkar, Bhushan N. Borse, Vinod D. Gupta, Kiran R. Phatangare, Vikas S. Patil, Prashant G. Umape, N. Sekar, Arabian Journal of Chemistry, 2016; 9: S1125-S1130.
  6. Synthesis and in-silico molecular docking simulation of 3-chloro-4-substituted-1-(2-(1H-benzimidazol-2-yl)phenyl)-azetid-2-ones as novel analgesic anti-inflammatory agent, Santosh S. Chhajed and Chandrashekhar D. Upasani, Arabian Journal of Chemistry, 2016; 9: S1779-S1785.
  7. New Mn(II), Ni(II), Cd(II), Pb(II) complexes with 2-methylbenzimidazole and other ligands. Synthesis, spectroscopic characterization, crystal structure, magnetic susceptibility and biological activity studies, Shayma A. Shaker, Hamid Khaledi, Shiau-Chuen Cheah, Hapipah Mohd Ali, Arabian Journal of Chemistry, 2016; 9: S1943-S1950.
  8. Mixed ligand complexation of some transition metal ions in solution and solid state: Spectral characterization, antimicrobial, antioxidant, DNA cleavage activities and molecular modeling, SuthaShobana, Jeyaprakash Dharmaraja, and Shanmugaperumal Selvaraj, Spectrochimica Acta Part A: Molecular and Biomolecular Spectroscopy, 2013; 107(15): 117-132.
  9. Zn(II), Cd(II) and Hg(II) complexes with 1-(*p*-methoxybenzyl)-2-(*p*-methoxyphenyl)benzimidazole: Syntheses, structures and luminescence, M. N. Manjunatha, Amol G. Dikundwar, K. R. Nagasundara, Polyhedron, 2011; 30(7): 1299-1304.
  10. Vibrational spectroscopic studies of metal(II) halide Benzimidazole, Ş. Yurdakul and M. Kurt, Journal of Molecular Structure, 2003; 650(1–3): 181-190.
  11. Synthesis and antitumor activity of novel benzimidazole-5-carboxylic acid derivatives and their transition metal complexes as topoisomerase II inhibitors, Shadia A. Galal<sup>a</sup>, Khaled H. Hegab, Ahmed M. Hashem and Nabil S. Youssef, European Journal of Medicinal Chemistry, December 2010; 45(12): 5685-5691.
  12. Coordination behavior of benzimidazole, 2-substituted benzimidazoles and benzothiazoles, towards transition metal ion, Fabiola Téllez, Horacio López-Sandoval, Silvia E. Castillo-Blum, and Noráh Barba-Behrens, ARKIVOC, 2008; 245-275.
  13. Synthesis and spectroscopic studies of bivalent transition metal complexes with Benzimidazole Derivative, Suman Malik, Supriya Das and Bharti Jain, RJPBCS, 2010; 1(3): 394-400.
  14. Synthesis and Biological Evaluation of Novel Benzimidazole Derivative with Aspirin as Potent Antimicrobial & Antifungal Agents Chavan B B, Chitte P D, Choudhary N P, Albhar K G, Dr. Hukkeri V I, IJSRR, 2012; 1(3): 22-30.
  15. Synthesis, Characterization and Evaluation of Novel N-(1H-Benzimidazol-2-Yl)-2-Isatinylidene-Hydrazinecarboxamide Derivatives as Anti-Inflammatory Agents, Gummadi Sridhar Babu, Nanam Rajani, Puttireddy S Malathy, Bethi Srinivas, Umasankar Kulandaivelu, Jupally Venkateshwar Rao, Der Pharma Chemica, 2010; 2(3): 196-204.
  16. Zinc(II) and Nickel(II) Benzoate Complexes from the Use of 1-methyl-4,5-diphenylimidazole Konstantina A. Kounavi, Manolis J. Manos, Anastasios J. Tasiopoulos, Spyros P. Perlepes, and Vassilios Nastopoulos, Bioinorganic Chemistry and Applications, 2010; 2010: 1-7.
  17. <sup>1</sup>H and <sup>13</sup>C-NMR spectra of condensed Benzimidazole and imidazobenzodiazepines, A. El kihel, E.M. Essassi, P. Bauchat, Arabian Journal of Chemistry, 2012; 5: 523-526.
  18. Synthesis, characterization and pharmacological screening of novel benzimidazole derivatives, Mohammad Shaharyar, Avijit Mazumder, Salahuddi, Ruchika Garg, Rishabh D Pandey, Arabian Journal of Chemistry, 2016; 9: S342-347.