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TETRACYCLINE COORDINATION COMPOUND AS ANTIMICROBIAL AGENT

Pranay Guru

Department of Engineering Chemistry, People's College of Research & Technology, Bhopal (M P), India

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ABSTRACT

The present paper deals with the microbial studies of the complex Co(II) & Cu(II) with antibiotic drug Tetracycline, a formula $\text{Co}(\text{C}_{22}\text{H}_{24}\text{N}_2\text{O}_8)\text{MoO}_4 \cdot 4\text{H}_2\text{O}$ and $\text{Cu}(\text{C}_{22}\text{H}_{24}\text{N}_2\text{O}_8)\text{MoO}_4 \cdot 3\text{H}_2\text{O}$ has been suggested on the basis of elemental analysis and molar conductance for the newly synthesized complex. The microbial studies of synthesized complex were studied on pathogenic bacteria using gram +ve (*Bacillus subtilis* and *Staphylococcus aureus*) and gram -ve (*Shigella flexneri*, *Salmonella typhosa*, *Escherichia coli*) and some fungi (*Aspergillus flavus*, *Fusarium oxysporum*, *Chrysosporium pannicale*, *Alternaria solani*, *Candida albicans*).

Correspondence to Author:

Dr. Pranay Guru

C- 20 , Siddhartha Lake City,
Raisen Road, Anand Nagar,
Bhopal (MP), India

INTRODUCTION: In continuation of the work being carried out in our laboratory on the metal molybdate with organic ligand ¹⁻⁵, the present communication describes microbial studies of Co(II) & Cu(II) with antibiotic drug tetracycline having molybdate as anion.

Experimental: Microbial studies of the synthesized complexes were performed at Department of Microbiology, Dr H S Gour University Sagar (MP) and Govt. Veterinary college Jabalpur (MP) using paper disc method Kathal *et al* ⁶, on the following pathogenic bacteria using gram +ve (*Bacillus subtilis* and *Staphylococcus aureus*) and gram -ve (*Shigella flexneri*, *Salmonella typhosa*, *Escherichia coli*) and some fungi (*Aspergillus flavus*, *Fusarium oxysporum*, *Chrysosporium pannicale*, *Alternaria solani*, *Candida albicans*).

RESULT & DISCUSSIONS: The synthesized complexes were screened for the antibacterial and antifungal activity using standard paper disc method ⁷⁻¹⁰ against gram positive bacterial viz (**table 1**). *Bacillus subtilis* and *Staphylococcus aureus* and gram negative bacteria viz. *Escherichia coli* and *Salmonella typhosa* and fungi *Aspergillus flavus*, *Alternaria solani*, *Candida albicans*, *Fusarium oxysporum* & *Chrysosporium pannicle*. In general all the tested complexes showed higher toxicity against bacterial and fungi under study.

TABLE 1: COMPLEXES OF DIFFERENT METALS WERE MARKED HAS S1, S2

S1-	Co (C ₂₂ H ₂₄ N ₂ O ₈) MoO ₄ 4H ₂ O
S2-	Cu (C ₂₂ H ₂₄ N ₂ O ₈) MoO ₄ 3H ₂ O

Antibacterial activity: From **table 2** it is concluded that complex S1 has shown maximum zone of inhibition against *Shigella flexneri* at the concentration of 0.1 M even at the concentration of 0.01 M it has shown good zone of inhibition in

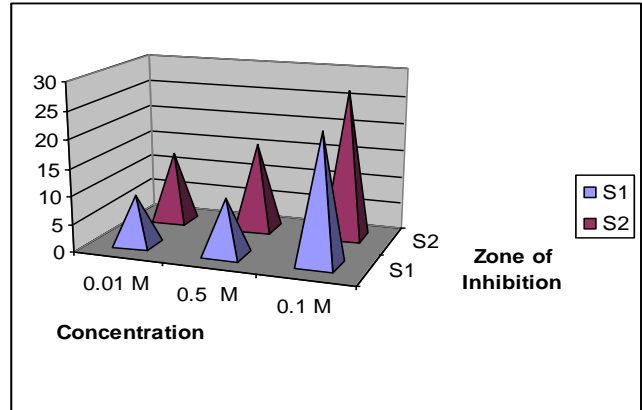
comparison to other tested complexes. Against *Salmonella typhosa* good antibacterial activity was observed against almost all the tested complexes. Complex S1 individually shown maximum zone of inhibition against this organism. Against *Escherichia-coli* maximum inhibitory effect were produced by complex S2. *Bacillus subtilis* was found to be more susceptible against complex S2 maximum zone of inhibition were recorded against these tested complexes. Complexes S2 found to be more active and shown higher zone of inhibition against *Staphylococcus aureus* in comparison to S1.

TABLE 2: ANTIBACTERIAL ACTIVITY OF SYNTHESIZED COMPLEXES

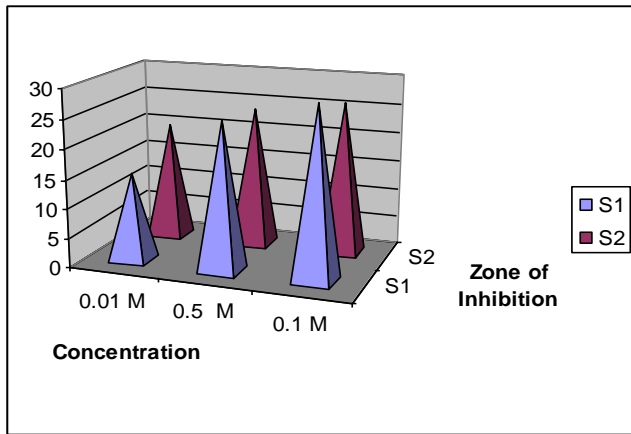
Bacteria	Concentration	Stain of Bacteria/ Zone of Inhibition (mm*)	
		S1	S2
<i>Shigella flexneri</i>	0.01 M	15	20
	0.5 M	25	24
	0.1 M	29	26
<i>Salmonella typhosa</i>	0.01 M	14	8
	0.5 M	24	19
	0.1 M	28	23
<i>Escherichia-coli</i>	0.01M	9	13
	0.5M	10	16
	0.1M	23	27
<i>Bacillus subtilis</i>	0.01M	9	12
	0.5 M	23	24
	0.1 M	25	27
<i>Staphylococcus aureus</i>	0.01 M	12	11
	0.5 M	18	15
	0.1 M	28	21

On comparing the anti-bacterial efficacy of these tested complexes, it is concluded that though most of the complexes reported satisfactory results for their antibacterial property but complexes S2 in particular gave promising results. From the above study it is observed that complex of Co(II) & Cu(II) with antibiotic drug Tetracycline

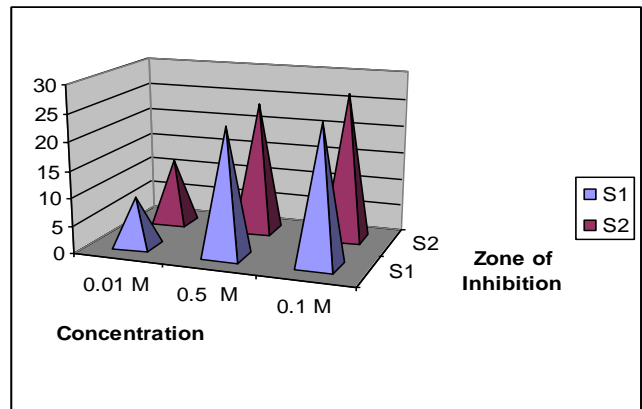
found to most active against the tested microorganisms. It is found that all the tested complexes exhibit good antibacterial activity at the concentration of 0.1 M and. it is interesting to note that inhibitory power of complexes decrease with the increase of their concentration. For the comparison of the antibacterial properties of these tested complexes against bacteria *Shigella flexneri*, *Salmonella typhosa*, *Escherichia- coli*, *Bacillus subtilis*, *Staphylococcus aureus* the zone of inhibition have been graphically represented in Graph 1 to 5.



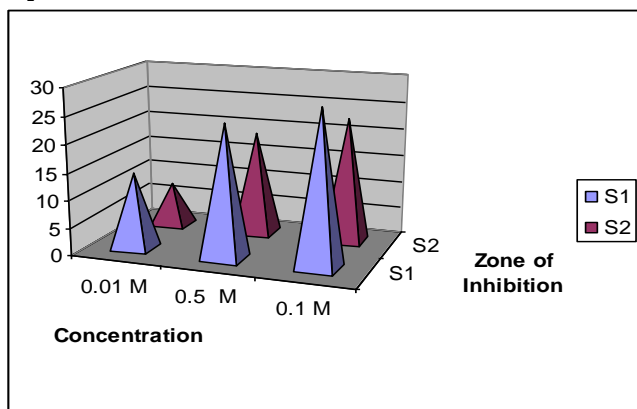
S1 = Ni (C₂₂ H₂₄ N₂ O₈) MoO₄ 3H₂O, S2 = Zn (C₂₂ H₂₄ N₂ O₈) MoO₄ 3H₂O
GRAPH 3: COMPARATIVE ANTIBACTERIAL ACTIVITY OF COMPLEXES AGAINST ESCEHRICHA-COLI



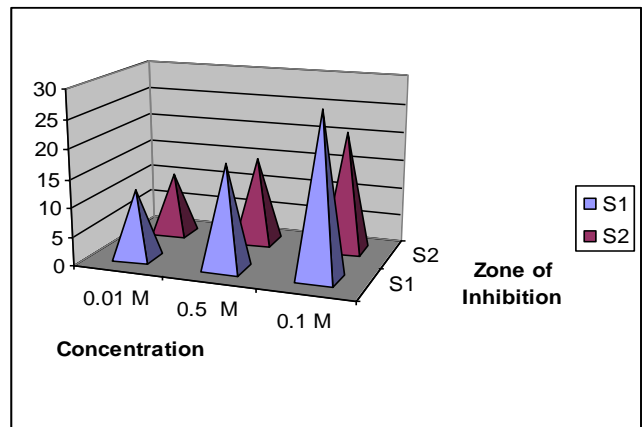
GRAPH 1: COMPARATIVE ANTIBACTERIAL ACTIVITY OF COMPLEXES AGAINST SHIGELLA FLEXNERI
 S1=Ni (C₂₂ H₂₄ N₂ O₈) MoO₄ 3H₂O, S2 = Zn (C₂₂ H₂₄ N₂ O₈) MoO₄ 3H₂O



GRAPH 4: COMPARATIVE ANTIBACTERIAL ACTIVITY OF COMPLEXES AGAINST BACILLUS SUBTILIS
 S1 = Ni (C₂₂ H₂₄ N₂ O₈) MoO₄ 3H₂O, S2 = Zn (C₂₂ H₂₄ N₂ O₈) MoO₄ 3H₂O



GRAPH 2: COMPARATIVE ANTIBACTERIAL ACTIVITY OF COMPLEXES AGAINST SALMONELLA TYPHOSA
 S1= Ni (C₂₂ H₂₄ N₂ O₈) MoO₄ 3H₂O, S2= Zn (C₂₂ H₂₄ N₂ O₈) MoO₄ 3H₂O



GRAPH 5: COMPARATIVE ANTIBACTERIAL ACTIVITY OF COMPLEXES AGAINST STAPHYLOCOCCUS AUREUS
 S1 = Ni (C₂₂ H₂₄ N₂ O₈) MoO₄ 3H₂O, S2 = Zn (C₂₂ H₂₄ N₂ O₈) MoO₄ 3H₂O

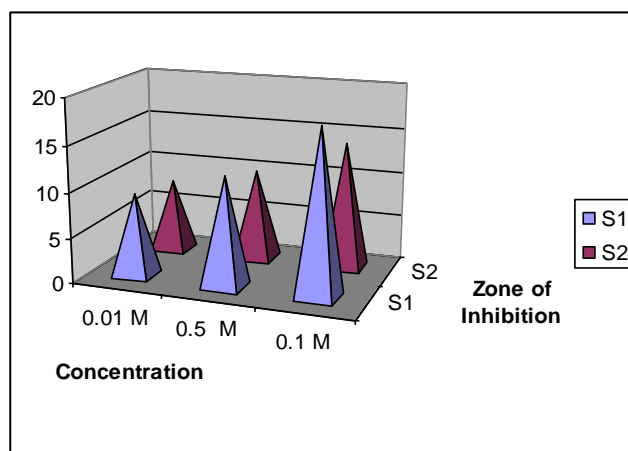
Anti fungal activity: Study of anti-fungal activity of complexes S1, S2 was carried out against selected five fungi namely *Aspergillus flavus*, *Candida albicans*, *Alternaria solani*, *Fusarium oxysporum* and *Chrysosporium pannicale*. at varying concentration of complexes 0.1M , 0.5 M and 0.01 M respectively and the result are recorded in terms of zone of inhibition which also includes the diameter of filter paper disc (6mm). From **table 3** it is observed that at the concentration of 0.1M of complex S1 shown maximum zone of inhibition was recorded against *Aspergillus flavus* similarly good inhibitory efficacy was also observed at the same concentration of complexes S2 against *Aspergillus flavus*.

TABLE 3: ANTIFUNGAL ACTIVITY OF SYNTHESIZED COMPLEXES

Fungi	Concentration	Stain of Fungi/ Zone of Inhibition (mm*)	
		S1	S2
<i>Aspergillus flavus</i>	0.01 M	9	8
	0.5 M	12	10
	0.1 M	18	14
<i>Candida albicans</i>	0.01 M	8	11
	0.5 M	10	17
	0.1 M	15	21
<i>Alternaria solani</i>	0.01M	11	9
	0.5M	15	12
	0.1M	23	18
<i>Fusarium oxysporum</i>	0.01M	8	7
	0.5 M	14	12
	0.1 M	26	22
<i>Chrysosporium pannicale</i>	0.01 M	9	7
	0.5 M	16	9
	0.1 M	23	14

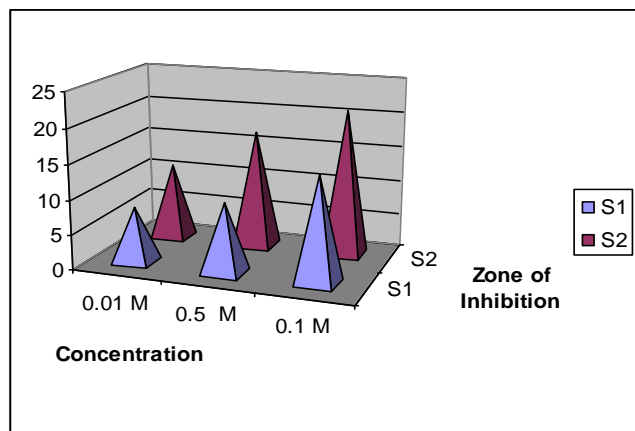
Against *Candida albicans* at the concentration of 0.1M complex S2 have shown maximum activity but similarly, considerable zone of inhibition were also recoded in case of complexes. It is evident from the result (Table 3) even at the concentration of 0.01 M all the complexes were found to be active against *Candida albicans*. Maximum zone of inhibition were recorded by all the complexes at the concentration of 0.1M against *Alternaria solani*. All the complexes at the concentration of 0.5 M have also given promising results.

The complex S2 at the concentration of 0.1 M produced maximum zone of inhibition against *Fusarium oxysporum*. Microorganism *Chrysosporium pannicale* was found susceptible against all the complexes tested at their concentration of 0.1M and 0.5 Complex S1 was found to posses good antifungal activity at 0.1 M concentration. For the comparison of the antifungal properties of these tested complexes against bacteria *Aspergillus flavus*, *Candida albicans*, *Alternaria solani*, *Fusarium oxysporum* and *Chrysosporium pannicale* the zone of inhibition have been graphically represented in Graph 6-10.



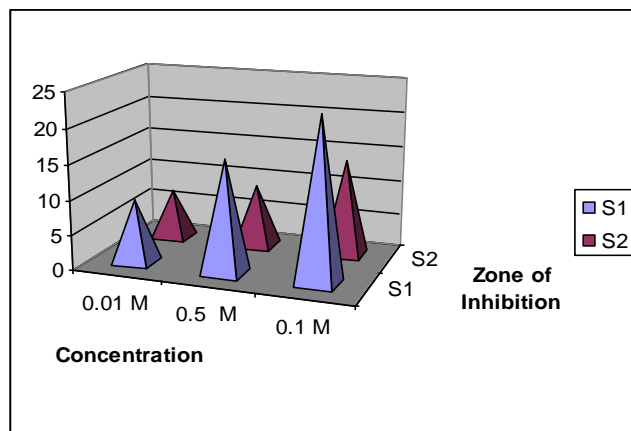
GRAPH 6: COMPARATIVE ANTIFUNGAL ACTIVITY OF COMPLEXES AGAINST ASPERGILLUS FLAVUS

S1 = Ni (C₂₂ H₂₄ N₂ O₈) MoO₄ 3H₂O, S2 = Zn (C₂₂ H₂₄ N₂ O₈) MoO₄ 3H₂O



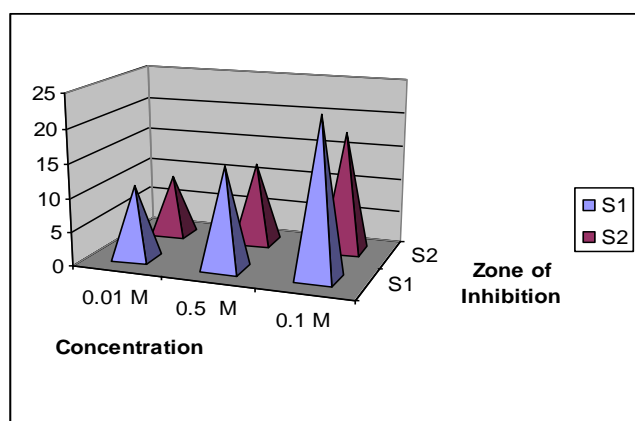
GRAPH 7: COMPARATIVE ANTIFUNGAL ACTIVITY OF COMPLEXES AGAINST *CANDIDA ALBICANS*

S1 = Ni (C₂₂ H₂₄ N₂ O₈) MoO₄ 3H₂O, S2 = Zn (C₂₂ H₂₄ N₂ O₈) MoO₄ 3H₂O



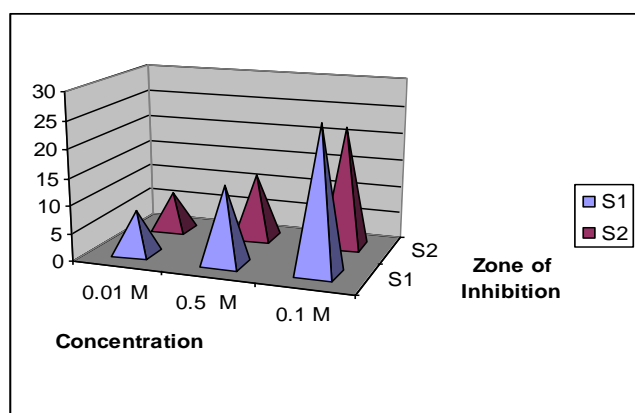
GRAPH 10: COMPARATIVE ANTIFUNGAL ACTIVITY OF COMPLEXES AGAINST *CHRYSOSPORIUM PANNICALE*

S1 = Ni (C₂₂ H₂₄ N₂ O₈) MoO₄ 3H₂O, S2 = Zn (C₂₂ H₂₄ N₂ O₈) MoO₄ 3H₂O



GRAPH 8: COMPARATIVE ANTIFUNGAL ACTIVITY OF COMPLEXES AGAINST *ALTERNARIA SOLANI*

S1 = Ni (C₂₂ H₂₄ N₂ O₈) MoO₄ 3H₂O, S2 = Zn (C₂₂ H₂₄ N₂ O₈) MoO₄ 3H₂O



GRAPH 9: COMPARATIVE ANTIFUNGAL ACTIVITY OF COMPLEXES AGAINST *FUSARIUM OXYSPORUM*

S1 = Ni (C₂₂ H₂₄ N₂ O₈) MoO₄ 3H₂O, S2 = Zn (C₂₂ H₂₄ N₂ O₈) MoO₄ 3H₂O

Antimicrobial properties of the original drug against selected microorganism were also compared. It could be observed that synthesized complex have shown promising result compared to commercial original drug Tetracycline.

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