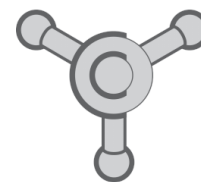




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The effect of plant extracts, *L. salicaria* and *S. pratensis* on the interaction between copper(II) complexes and biomolecules



Lythrum salicaria

Andrija D. Gigić¹, Ana S. Kesić², Jovana P. Bugarinović¹, Nikola Z. Srećković¹, Vladimir B. Mihailović¹, Jovana V. Bogojeski¹

¹ University of Kragujevac, Faculty of Science, Kragujevac, Serbia

² University of Kragujevac, Institute for Information Technologies Kragujevac, Department of Science, Kragujevac, Serbia

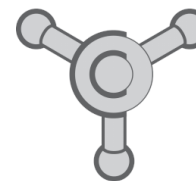


Salvia pratensis



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Introduction

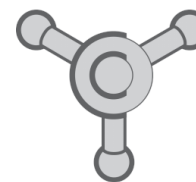
Copper complexes have demonstrated remarkable and selective biological activity, attracting attention as promising anticancer metallodrugs. Their potential arises from their complex mechanisms of action, where they interact with various cellular components and influence critical processes related to cell survival and proliferation.¹

1. L. A. Alfonso-Herrera, J. A. Aragón-Muriel, et al., ChemMedChem 2022, 17 (18)



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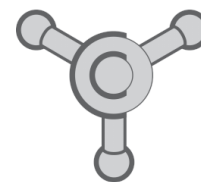
Ferrocene derivatives are valued for their reversible redox properties, high stability, and lipophilicity, which facilitate interactions with biomolecules such as DNA and proteins. When incorporated into Schiff base ligands, ferrocenyl groups can synergistically enhance both the anticancer and antimicrobial activity of copper complexes, forming multifunctional scaffolds with strong therapeutic promise.²

1. L. A. Alfonso-Herrera, J. A. Aragón-Muriel, et al., ChemMedChem 2022, 17 (18)
2. S. Peter, E. Morifi, et al., Pharmaceutics 2025, 17 (6), 722



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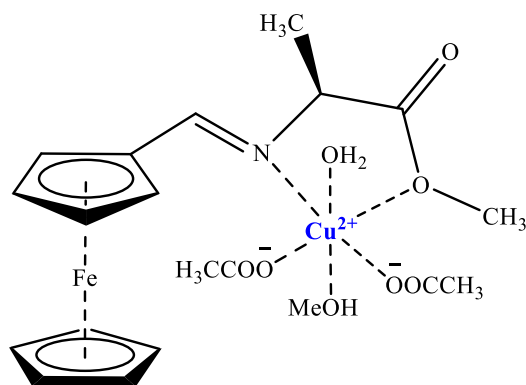
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Combining copper complexes with active ligands and **natural compounds** offers a unique strategy to exploit potential synergistic effects, where the pharmacological potential of phytochemicals complements the redox and coordination chemistry of the metal complexes.³

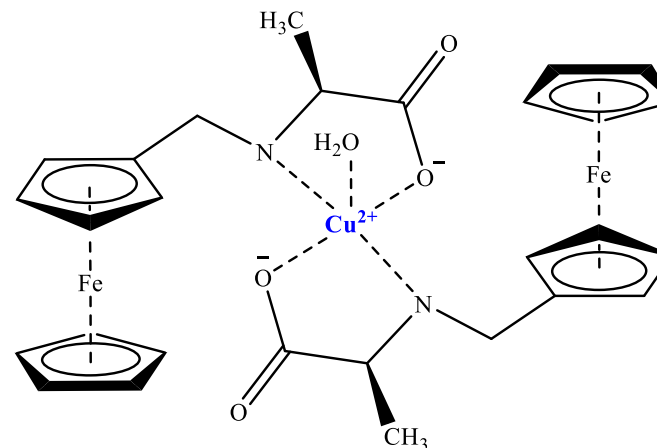
1. L. A. Alfonso-Herrera, J. A. Aragón-Muriel, et al., ChemMedChem 2022, 17 (18)
2. S. Peter, E. Morifi, et al., Pharmaceutics 2025, 17 (6), 722
3. N. Srećković, J. Katanić Stanković, S. Matić, N. Mihailović, P. Imbimbo, DM Monti, V. Mihailović, Ind. Crops Prod.. 2020, 155, 112781.

Introduction

Copper(II) complexes



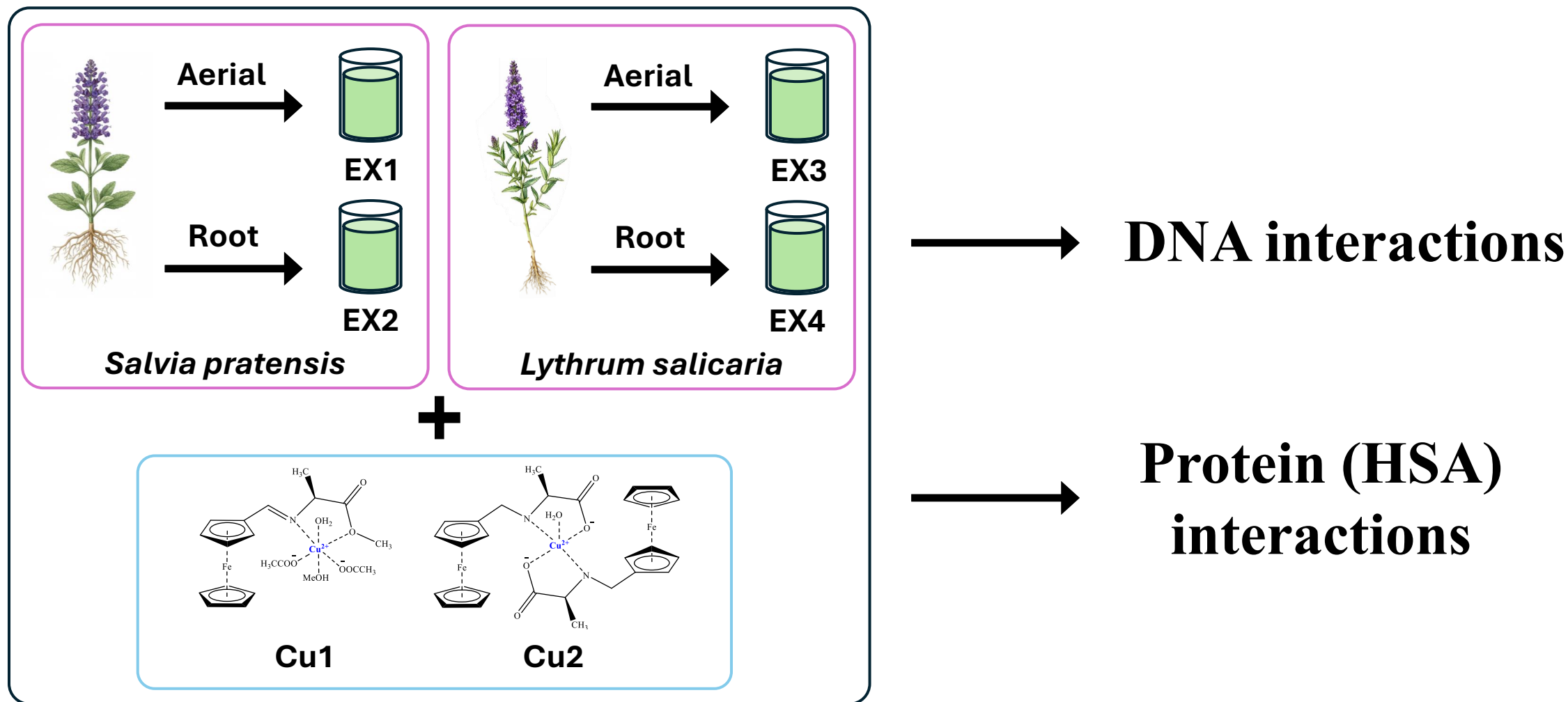
Cu1



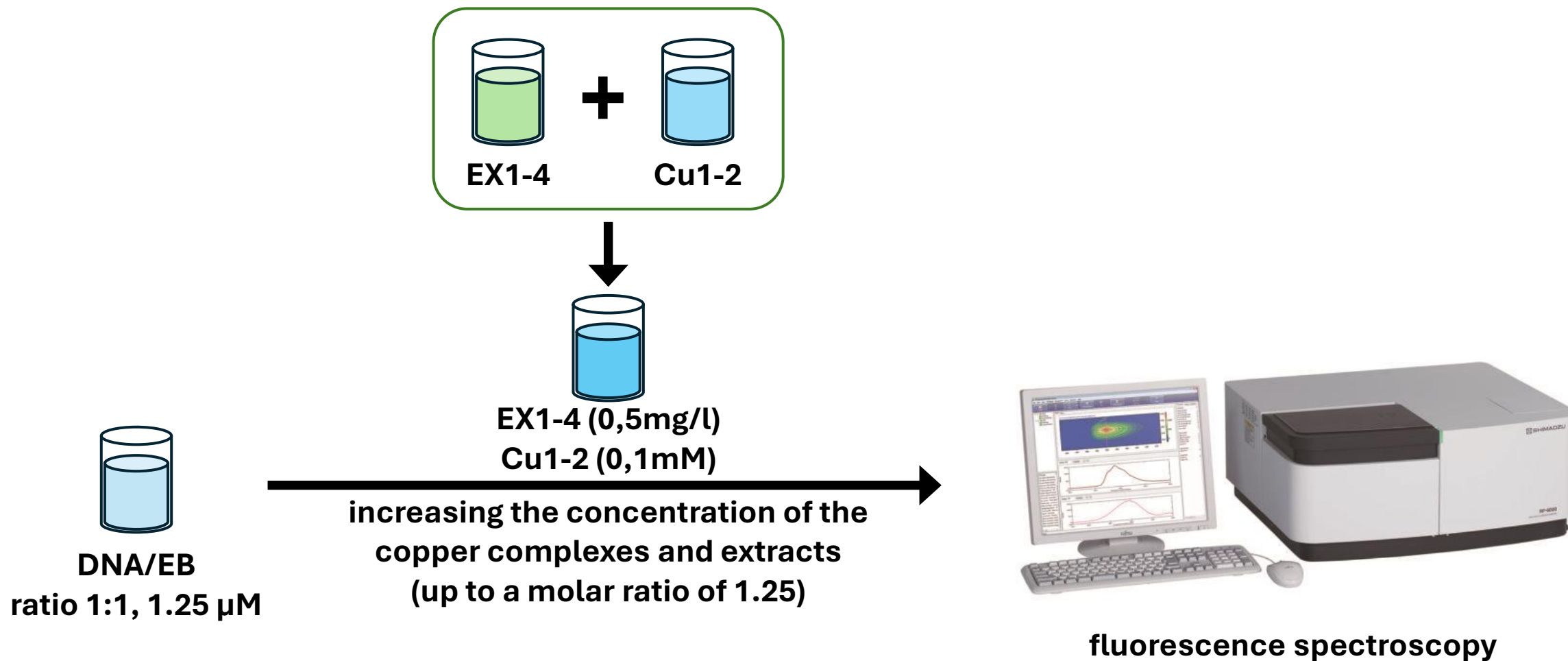
Cu2

Characterization: UV-Vis, IR, and mass spectroscopies, while single crystals of the **Cu2** complex were also obtained.

Introduction



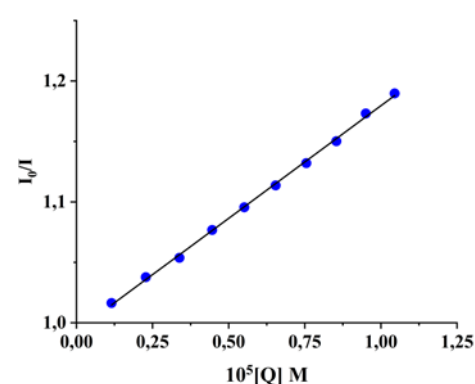
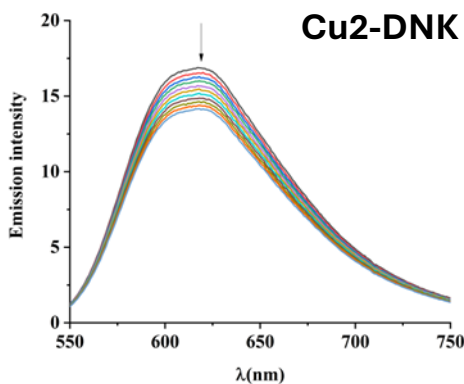
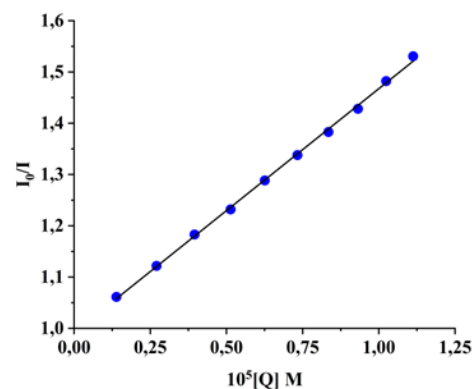
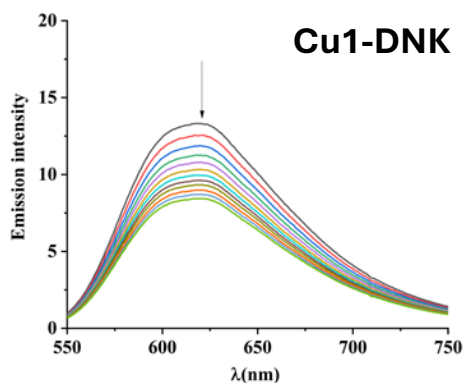
DNA interactions



DNA interactions



Cu1-2



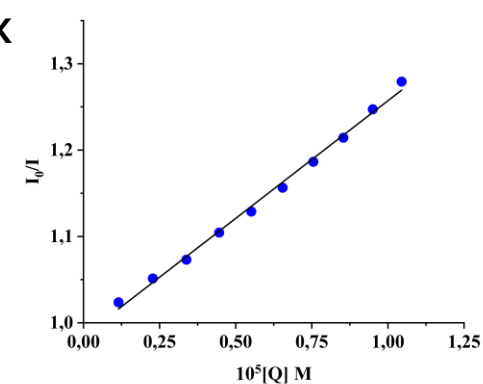
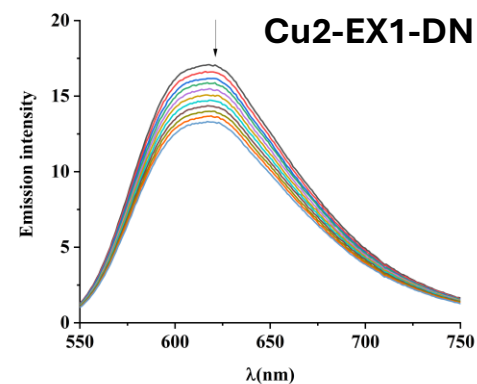
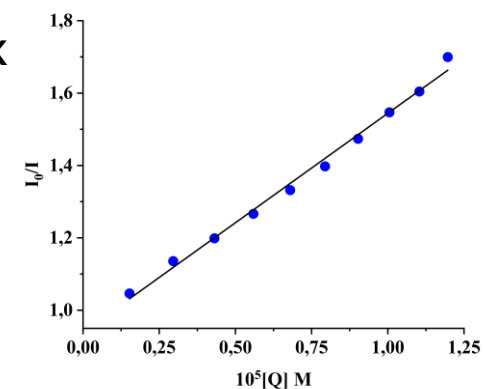
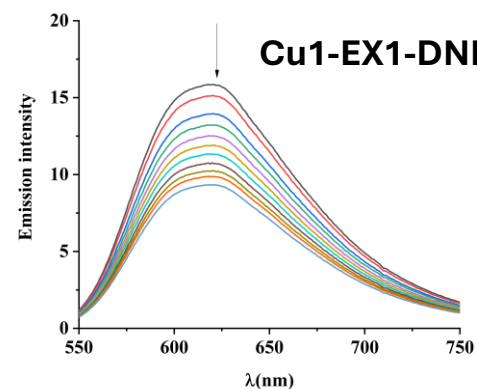
EX1



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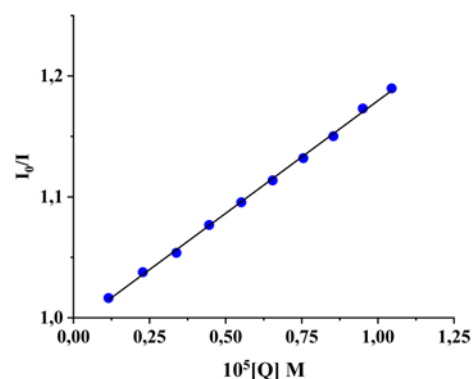
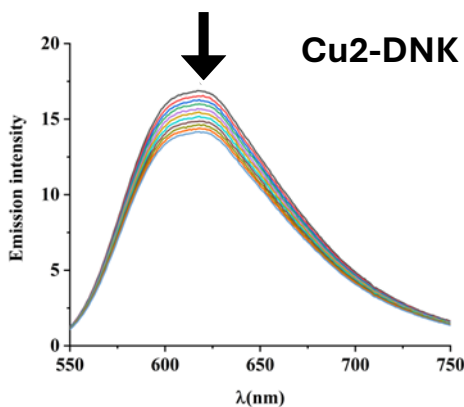
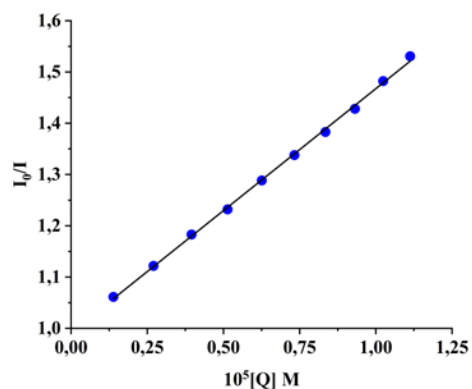
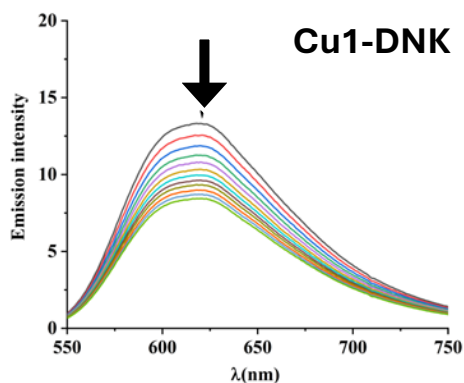
Cu1-2



DNA interactions



Cu1-2



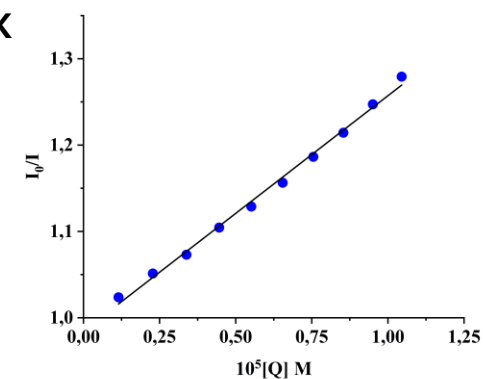
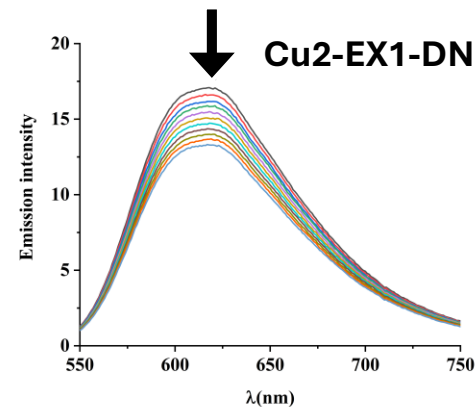
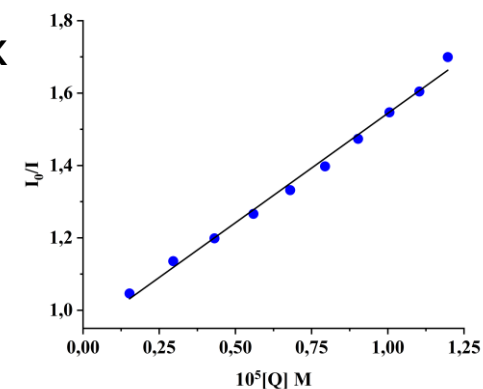
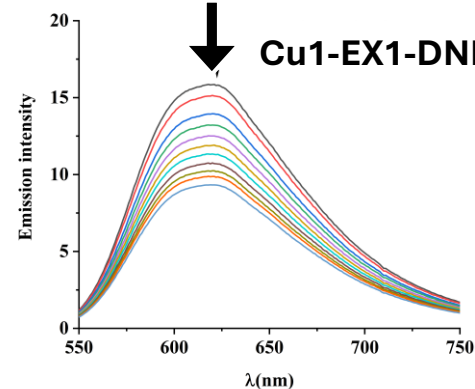
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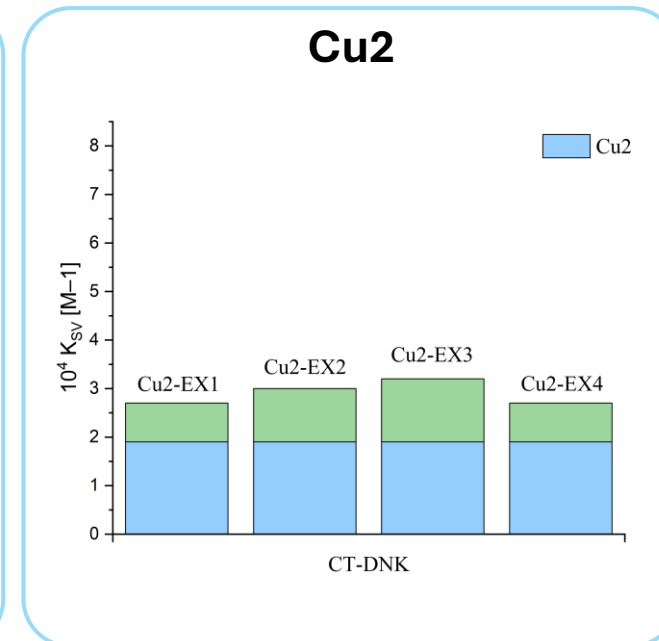
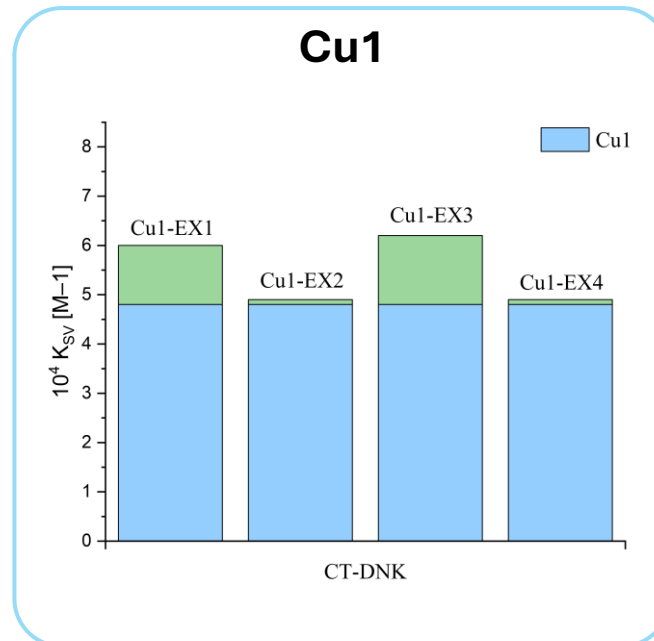


Cu1-2



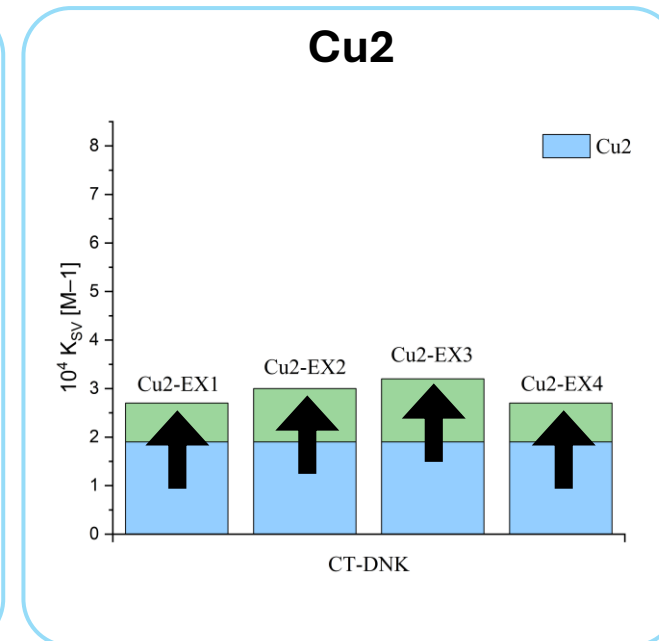
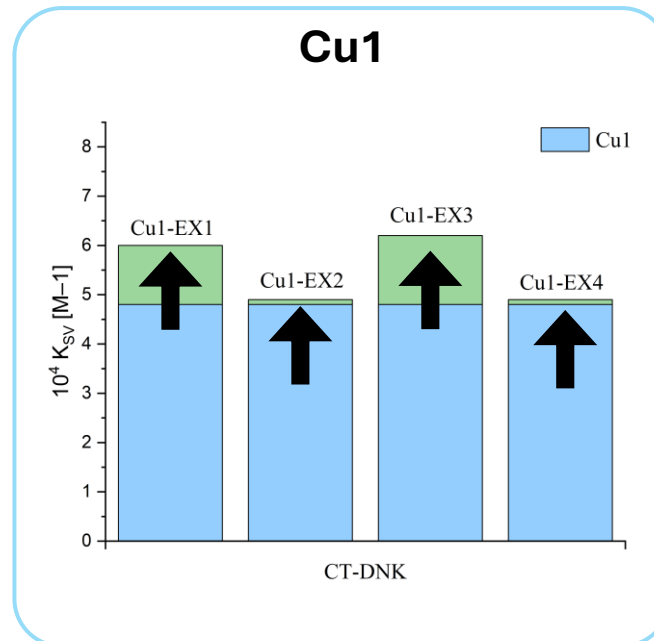
DNA interactions

DNK	Cu1	Cu2
	$10^4 K_{SV} [M^{-1}]$	$10^4 K_{SV} [M^{-1}]$
	4.8 ± 0.1	1.9 ± 0.1
EX1	6.0 ± 0.1	2.7 ± 0.1
EX2	4.9 ± 0.1	3.0 ± 0.1
EX3	6.2 ± 0.1	3.2 ± 0.1
EX4	4.9 ± 0.1	2.7 ± 0.1

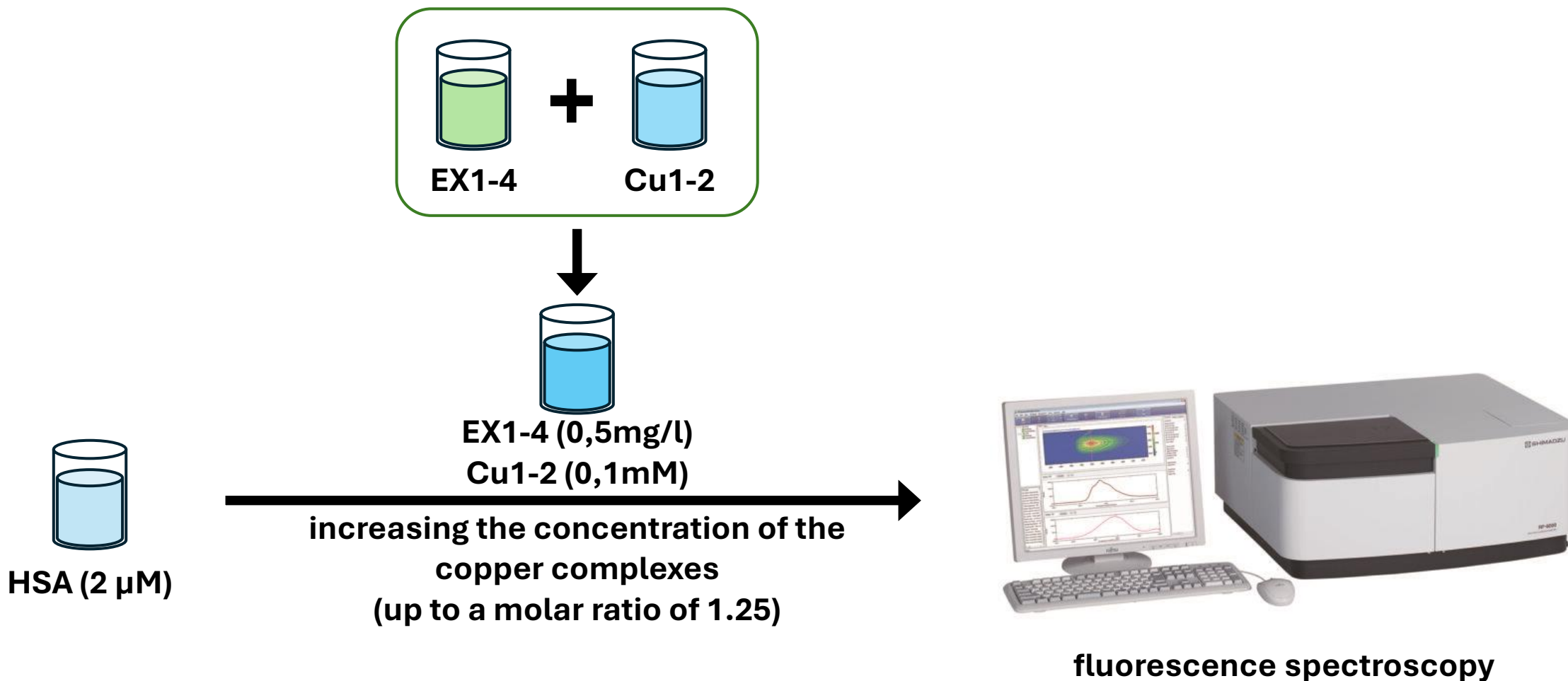


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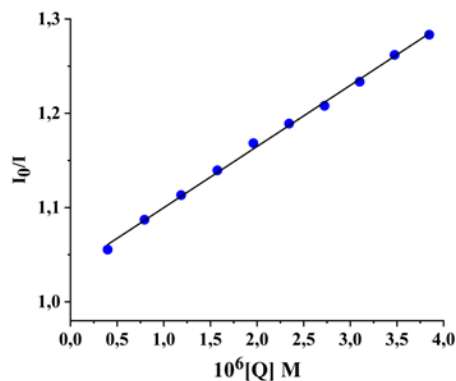
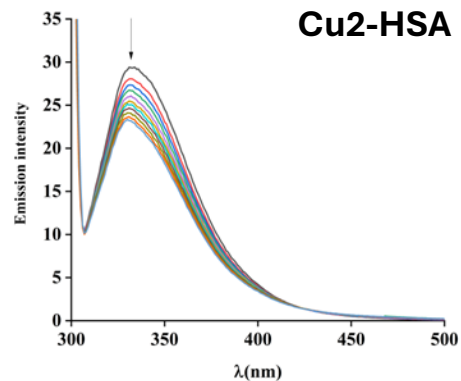
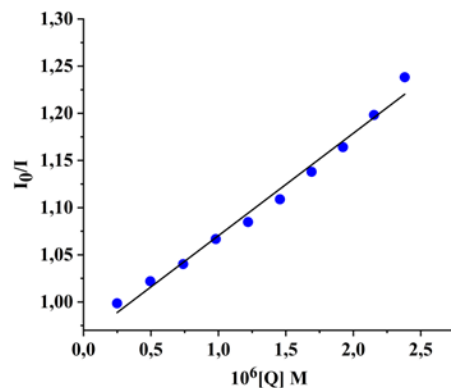
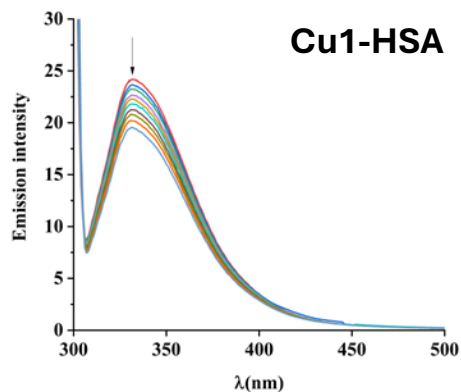
Protein (HSA) interactions



Protein (HSA) interactions



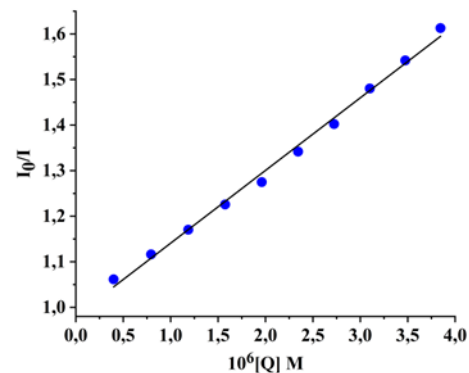
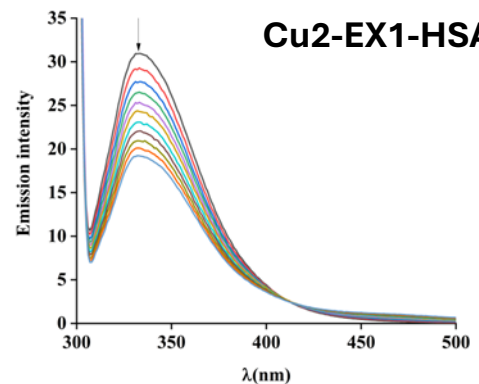
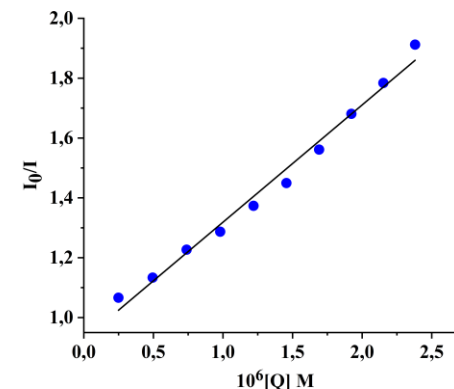
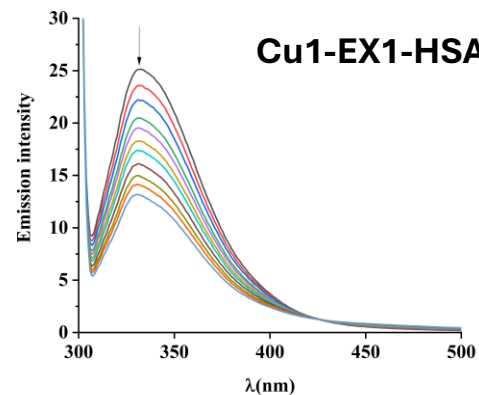
Cu1-2



EX1



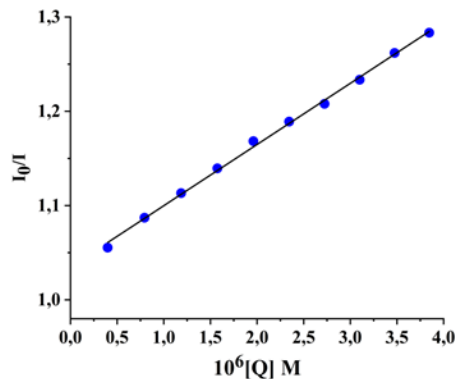
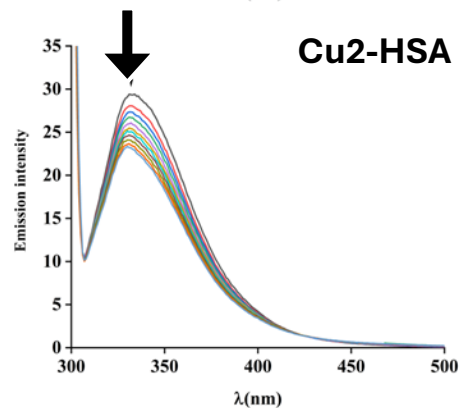
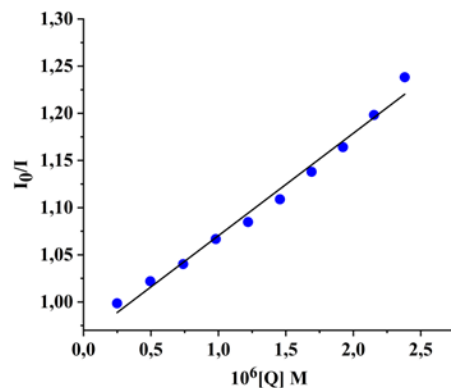
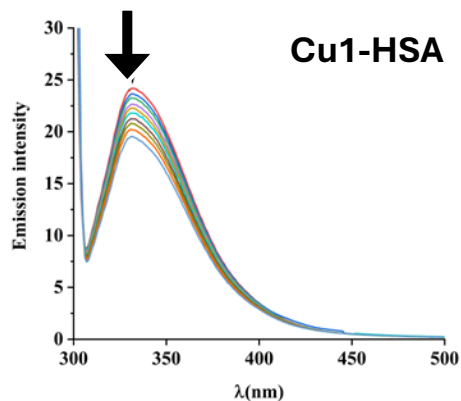
Cu1-2



Protein (HSA) interactions



Cu1-2



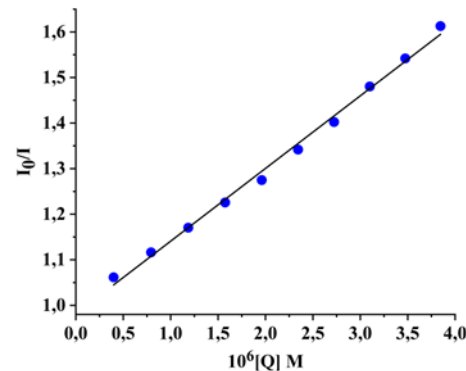
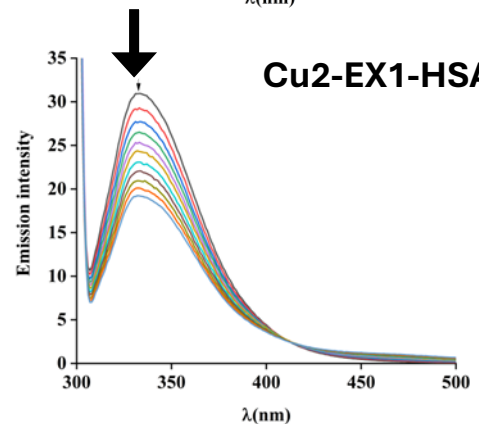
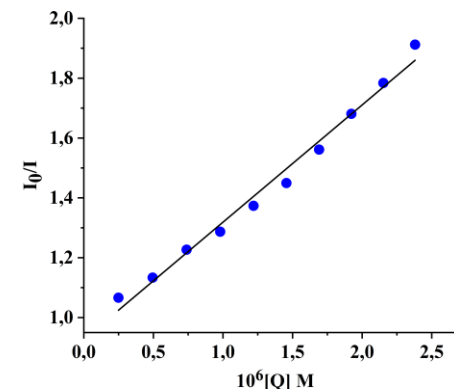
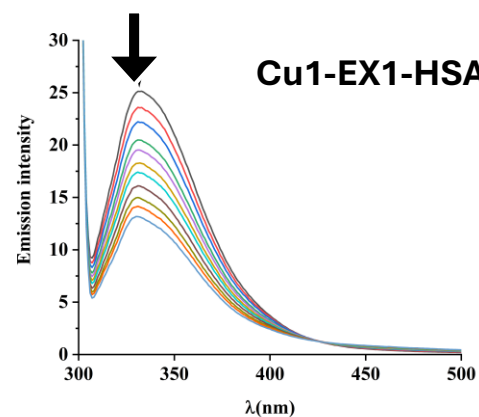
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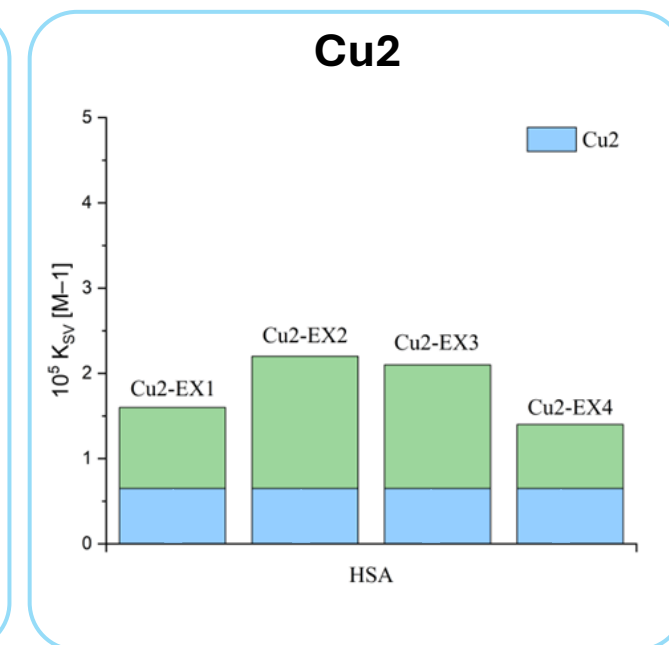
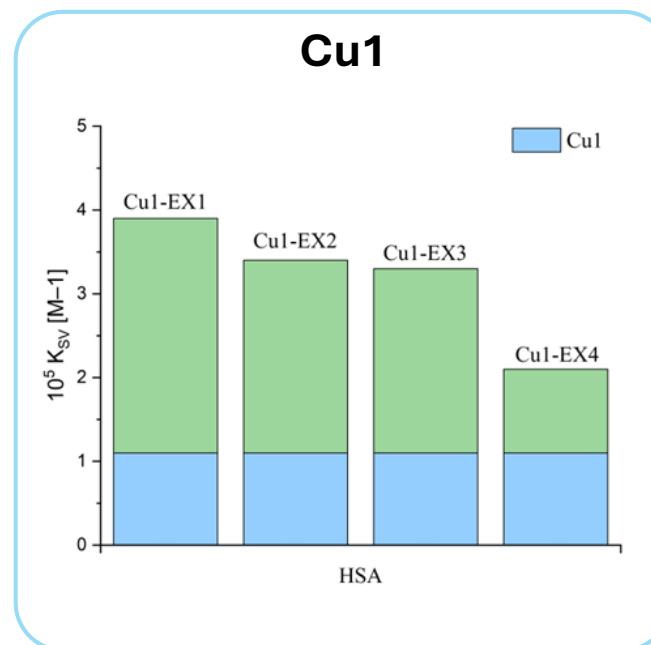


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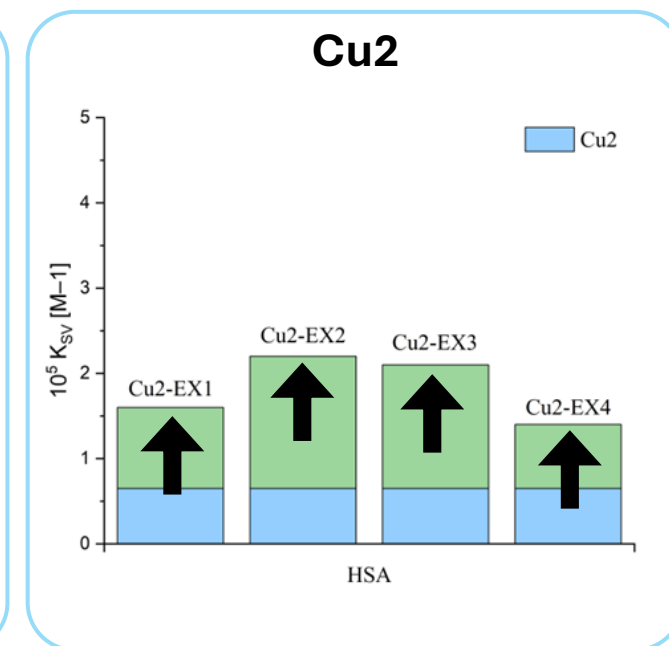
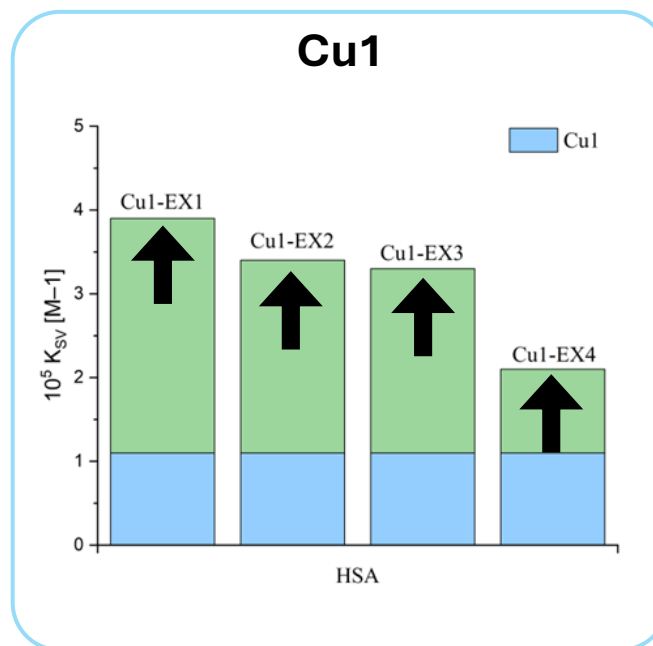
Protein (HSA) interactions

HSA	Cu1	Cu2
	$10^5 K_{sv} [M^{-1}]$	$10^5 K_{sv} [M^{-1}]$
	1.1 ± 0.1	0.65 ± 0.1
EX1	3.9 ± 0.1	1.6 ± 0.1
EX2	3.4 ± 0.1	2.2 ± 0.1
EX3	3.3 ± 0.1	2.1 ± 0.1
EX4	2.1 ± 0.1	1.4 ± 0.1



Protein (HSA) interactions

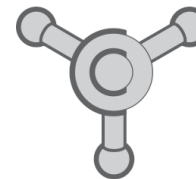
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Discussion and Future Research

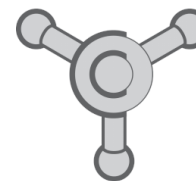
Structure matters

Cu1, the Schiff base, consistently binds stronger than **Cu2** because its geometry allows better access to DNA and protein binding pockets.



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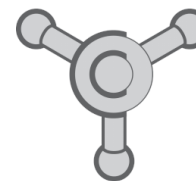
Synergy matters

Plant extracts significantly enhance the binding of both complexes, pointing to a cooperative effect between synthetic complexes and natural molecules.



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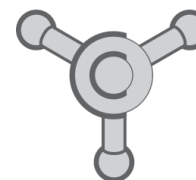
Implications

These findings suggest that combining synthetic metal complexes with natural products could be a promising approach for designing compounds with improved biological activity, potentially relevant for therapeutic applications.



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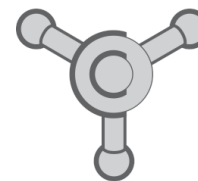
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Building on these promising results, several research directions are worth pursuing: the synthesis of similar ligands and complexation with other metal ions, as well as the investigation of antimicrobial and antitumor activities.



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Thank you for your attention.



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Vladimir B. Mihailović¹



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**Special thanks to all my colleagues and mentors
for their guidance, collaboration, and encouragement during this work.**