

Effect of Tyrosol Functionalized Gold Nanocomposite on Biofilm Milieu and Wound Healing

Tara Chand Yadav¹,

Payal Gupta², Ramasare Prasad³, Vikas Pruthi³, Akash Bachhuka¹, Lluís F. Marsal¹

¹Department Eng. Electronics, Electrical and Automation, Rovira i Virgili University, Tarragona, Spain

²Department of Biotechnology, Graphic Era University, Dehradun, India.

³Department of Biosciences & Bioengineering, Indian Institute of Technology Roorkee, Roorkee, Uttarakhand, India

tarachand.iitr@gmail.com

Abstract

Candidiasis, a fungal infection caused by opportunistic pathogen *Candida* which range from superficial to deep invasive manifestations, and is of significant concern in immunocompromised individuals [1]. Moreover, the fungal infections provide conditioned space to bacterial pathogens for colonization and results in mixed/ polymicrobial surface adhered community called polymicrobial biofilm. These communities are particularly responsible for emergence of multidrug-resistant strains. Thus, posed formidable challenges for clinicians in effectively managing these infections using conventional antimicrobial therapies. Hence, we synthesized chitosan mediated gold nanocomposite of tyrosol, for their efficacy against *Candida albicans*, *Staphylococcus aureus*, *Pseudomonas aeruginosa* and their mixed communities. Compared to tyrosol alone, Chi-TY-AuNP's nanocomposite demonstrated excellent inhibition potential on both planktonic and sessile growth of *C. albicans*, *S. aureus*, *P. aeruginosa*, *C. albicans* + *S. aureus*, and *C. albicans* + *P. aeruginosa* at lower concentrations. At the highest concentration tested (20 µg/mL), the drug conjugated nanoparticle has eradicated nearly 70% of the mature/ old polymicrobial biofilm structure while completely inhibited the construction of community. To ensure the suitability of the Chi-TY-AuNP's in treatment of non-healing wounds in diabetic patients, the cytocompatibility and wound healing was also undertaken [2].

Chi-TY-AuNP's treatment revealed good cytocompatibility and decreased ROS levels in NIH-3T3 cell lines. Developed 1% and 2% (w/w) Chi-TY-AuNP's-Carbopol@934 formulations exhibited excellent rheological properties suitable for topical applications, supported by the nontoxic nature of developed gels in the dermal toxicity study in experimental rats. Findings of *in-vivo* wound healing study conclusively demonstrated the re-epithelialization of the wounded tissues with no residual scar. Increased collagen and fibroblast

depositions, normal cellular integrity and neoangiogenesis were evident by the H&E and Masson's trichrome stained histopathological micrographs of the treated animal's skin tissues. The wound-healing effect mediated by Chi-TY-AuNP's-Carbopol@934 can be attributed to the additive antimicrobial and anti-inflammatory effects of tyrosol and chitosan in lipid peroxidation and oxidative damage minimizing activity [2]. Also, tyrosol being a polyphenolic compound, may act as a regulator at multiple phases of the wound healing process via preventing inflammation and secondary infection and augmenting collagen synthesis [3]. Collectively, *in-vivo* wound healing study demonstrated the reepithelialization of the wounded tissues and increased proliferation of fibroblast cells, collagen fibers and neoangiogenesis [2,4].

References

- [1] Yadav, T. C., Gupta, P., Saini, S., Mohiyuddin, S., Pruthi, V., & Prasad, R., ACS omega, 7(10), (2022), 8350-8363.
- [2] Yadav, T. C., Gupta, P., Pruthi, V., & Prasad, R. (2023) *under communication*
- [3] Yadav, T. C., Kumar, N., Raj, U., Goel, N., Vardawaj, P. K., Prasad, R., & Pruthi, V. Journal of Biomolecular Structure and Dynamics. 38:2, (2019), 382-397.
- [4] Raghuvanshi, N., Kumari, P., Srivastava, A. K., Vashisth, P., Yadav, T. C., Prasad, R., & Pruthi, V. Materials Science and Engineering: C, 80, (2017), 252-262.

Figures

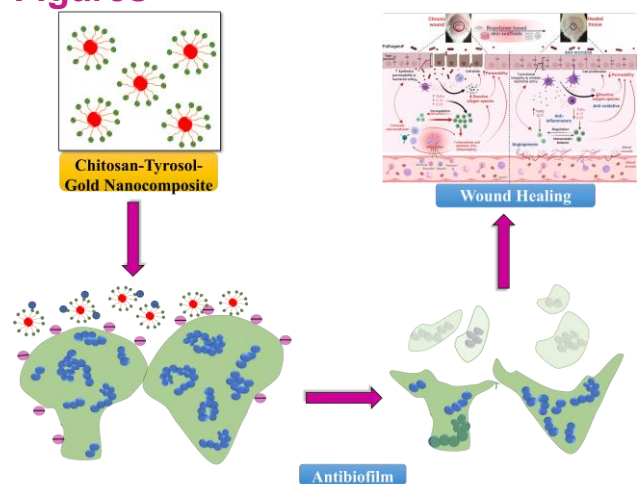


Figure 1. Chi-TY-AuNP's mediated Antibiofilm and Wound Healing activity.