

# Protocatechualdehyde-Enhanced Injectable Oxidized Pectin–Gelatin Hydrogels for Advanced Wound Healing and Tissue Regeneration

Nazlıcan Demircan<sup>a\*</sup>, Umut Gökay Ünal<sup>b</sup>, Somayeh Hormaty<sup>c</sup>, Gülşah Torkay<sup>c</sup>, Ayça Bal-Öztürk<sup>c</sup>, Banu Kocağa<sup>a,b</sup>

<sup>a</sup> Üsküdar University, Faculty of Engineering and Natural Sciences, Bioengineering, İstanbul, 34000, Turkey

<sup>b</sup> İstanbul Technical University, Faculty of Chemical and Metallurgy, Chemical Engineering, İstanbul, 34000, Turkey

<sup>c</sup> İstinye University, Faculty of Engineering and Natural Sciences, Biomaterials Stem Cell and Tissue Engineering, İstanbul, 34000, Turkey

\*E-mail: demircannazlican@gmail.com



## BACKGROUND

- Hydrogels with dynamic **Schiff base linkages** offer biocompatibility, self-healing, and promote cell adhesion, making them ideal for biomedical use.
- This study presents a **dialdehyde-functionalized pectin (OP)** - gelatin hydrogel, where both polymers support bioactive delivery, tissue regeneration, and tissue adhesion. **Protocatechualdehyde (PA)** reinforces the system with antioxidant, antibacterial, and anti-inflammatory effects .
- **Procaine (PC)** is incorporated for localized and **sustained release**, contributing both anesthetic and antioxidant functions [1] .

## OBJECTIVES

This study aims to develop an **injectable, self-healing, and tissue-adhesive scaffold** based on Schiff base linkages among OP, gelatin, and PA, aiming to achieve high **antioxidant capacity**, controlled PC release, and promotion of **angiogenesis**.

## METHODOLOGY

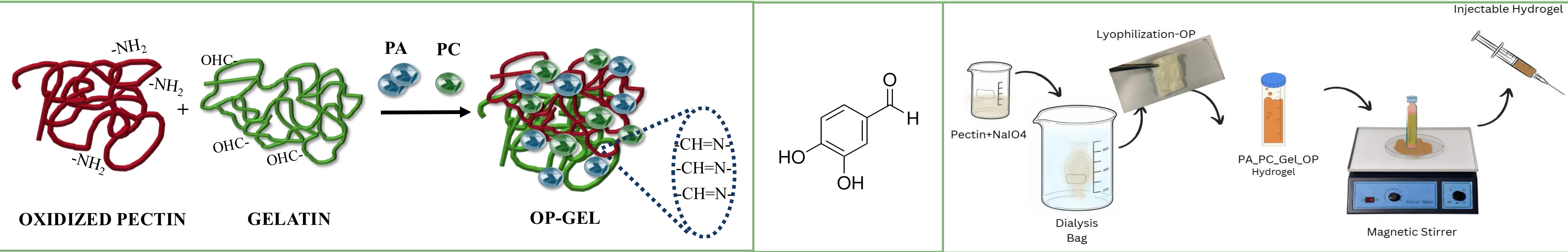


Figure 1. Oxidized pectin-gelatin crosslink formation and Protocatechualdehyde

Figure 2. Protocatechualdehyde

Figure 3. Schematic representation of the experimental procedure

## RESULTS

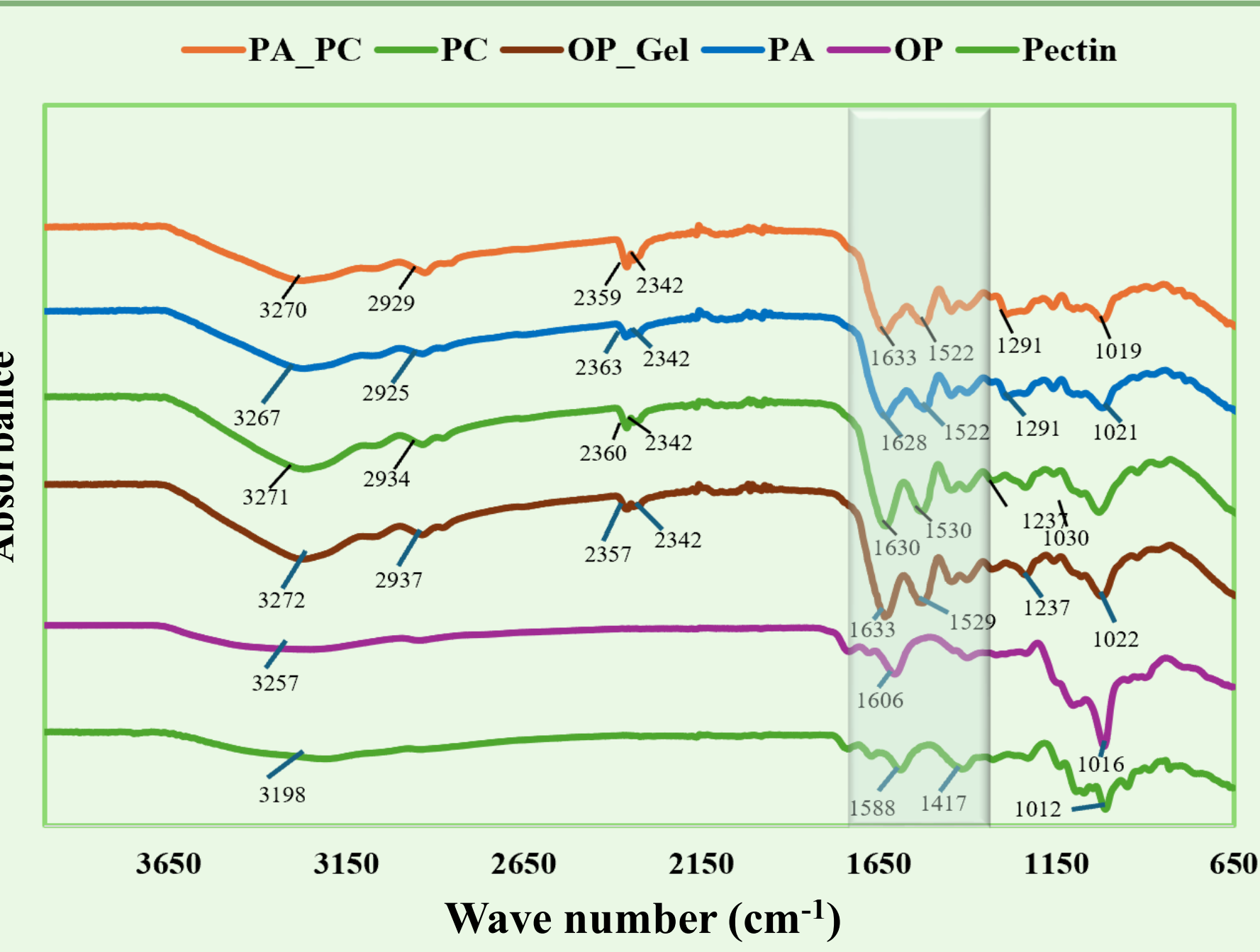


Figure 4. FTIR spectrum

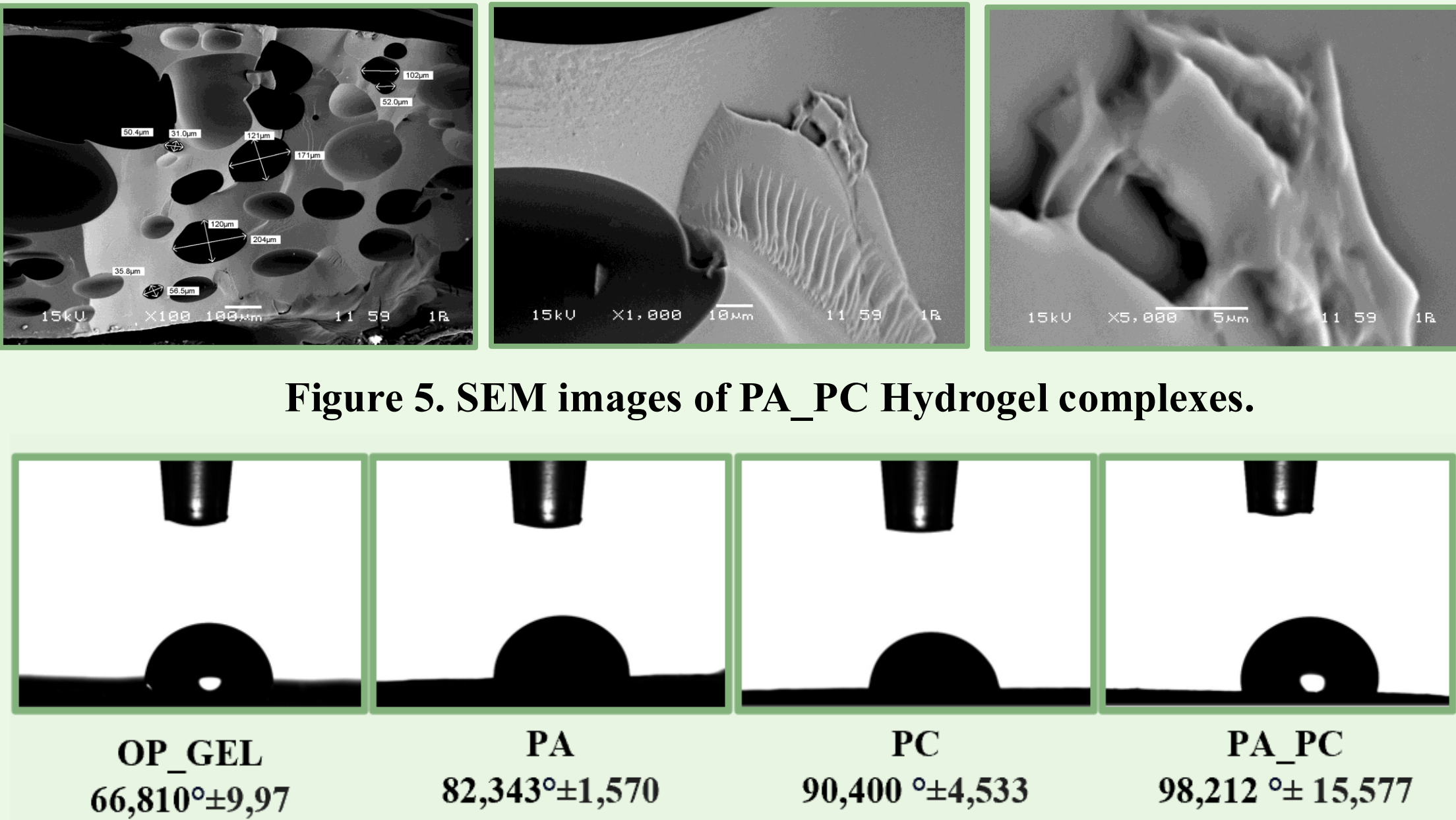


Figure 5. SEM images of PA\_PC Hydrogel complexes.

Figure 6. Contact Angle Measurement

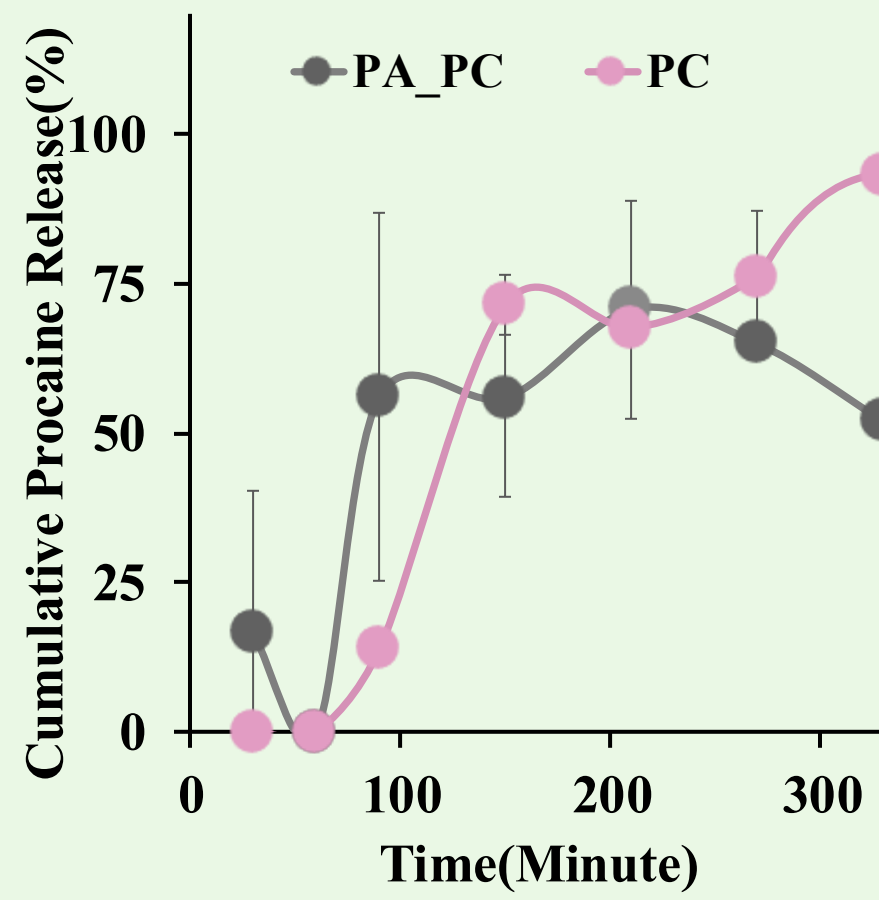


Figure 7. Procaine Release

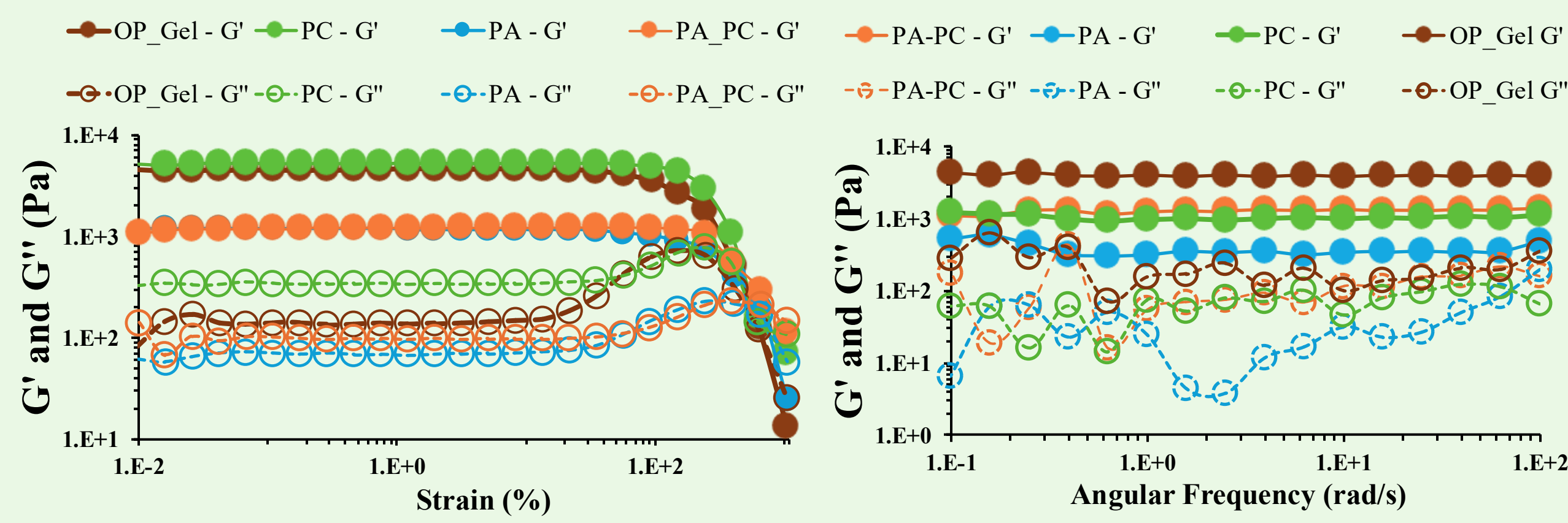


Figure 8. Amplitude Sweep

Figure 9. Frequency Sweep

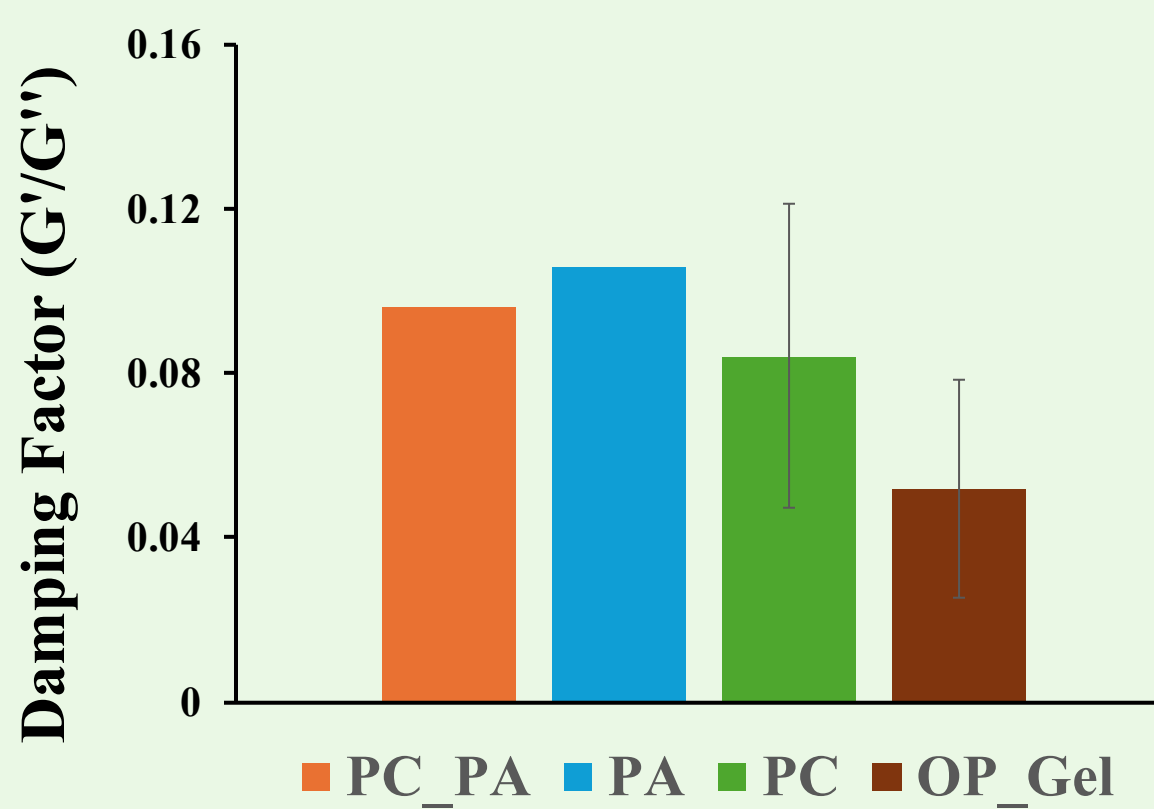


Figure 10. Damping Factor

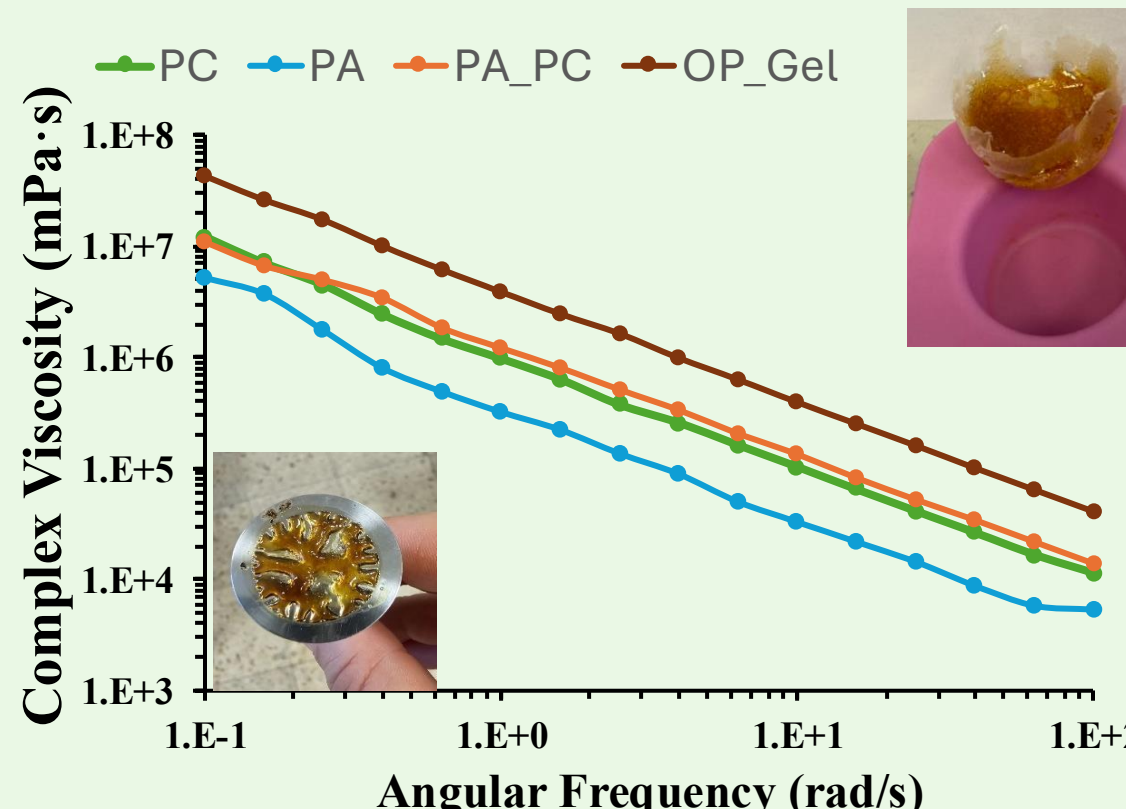


Figure 11. Complex Viscosity

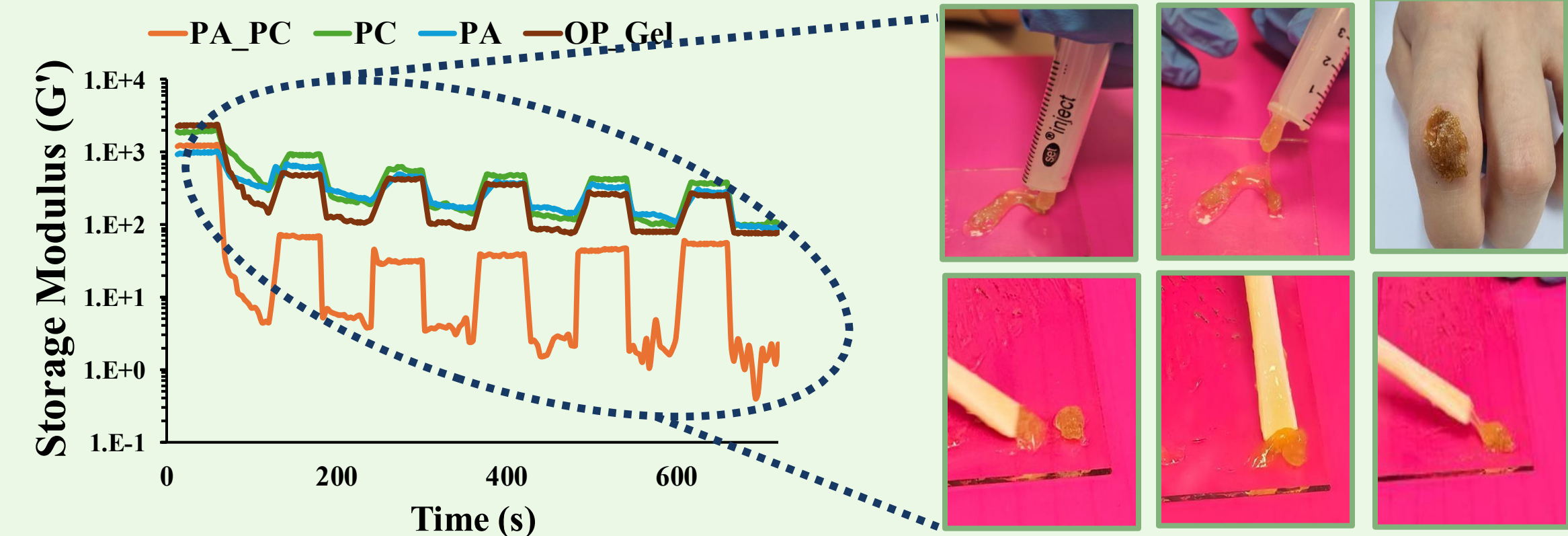


Figure 12. Self Healing

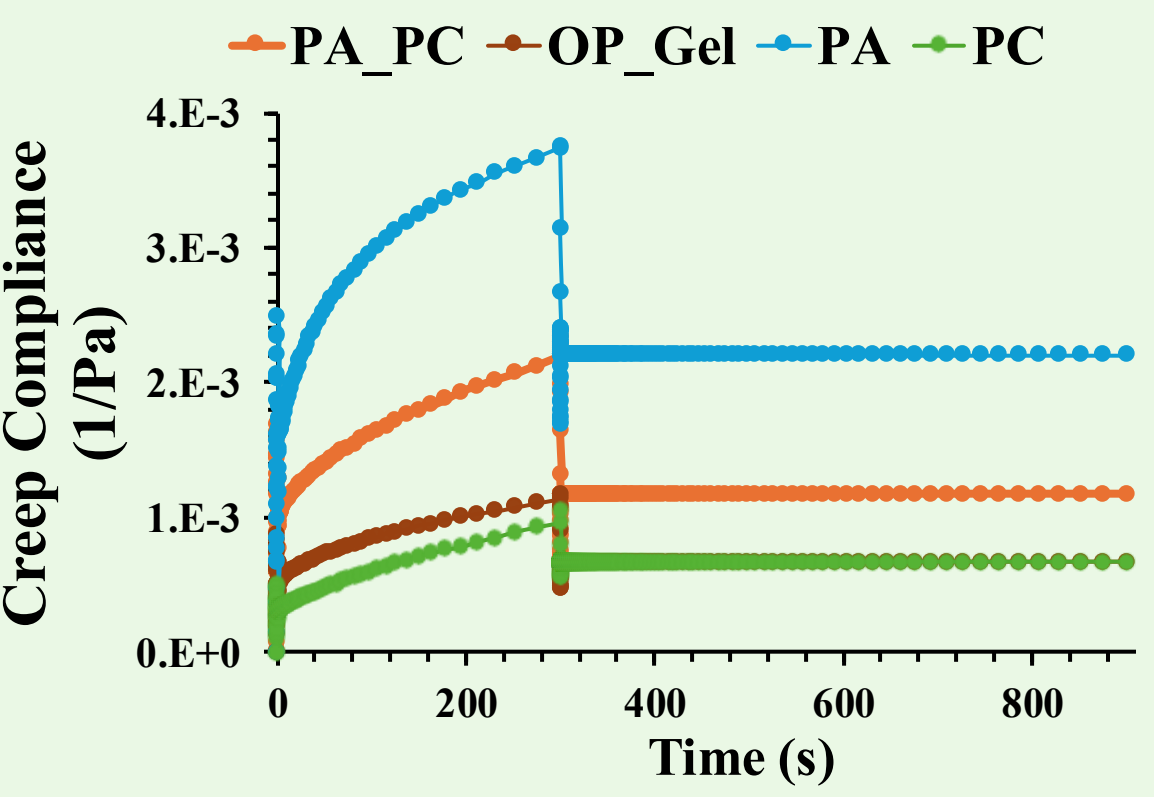


Figure 13. Creep

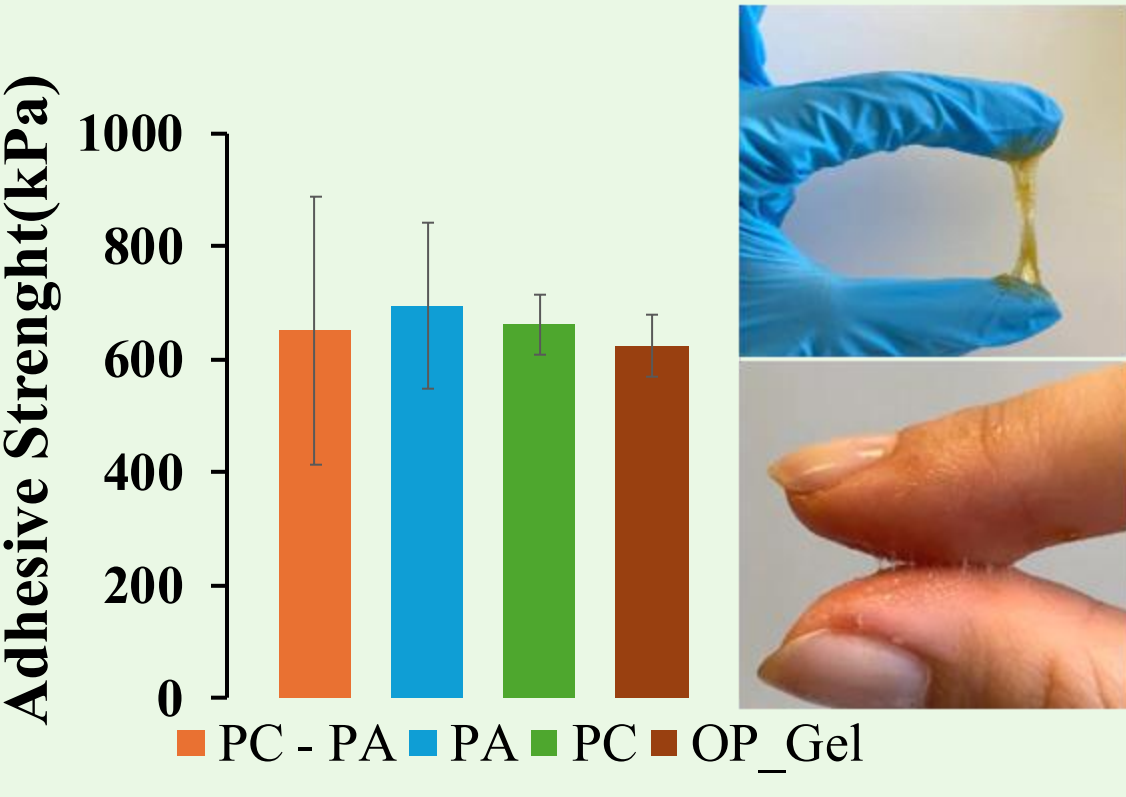


Figure 14. Adhesive strength

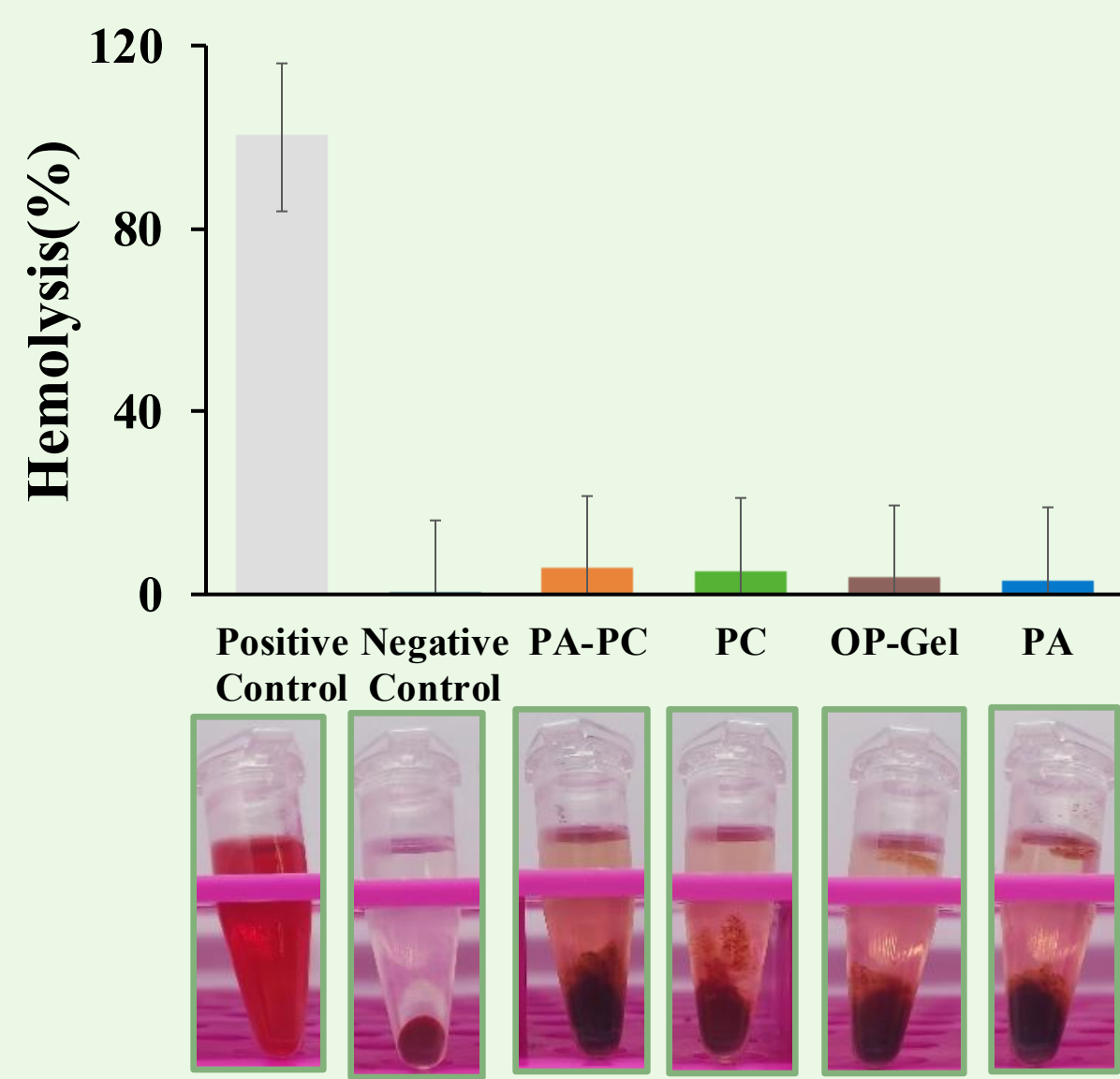


Figure 15. Hemolysis Analysis

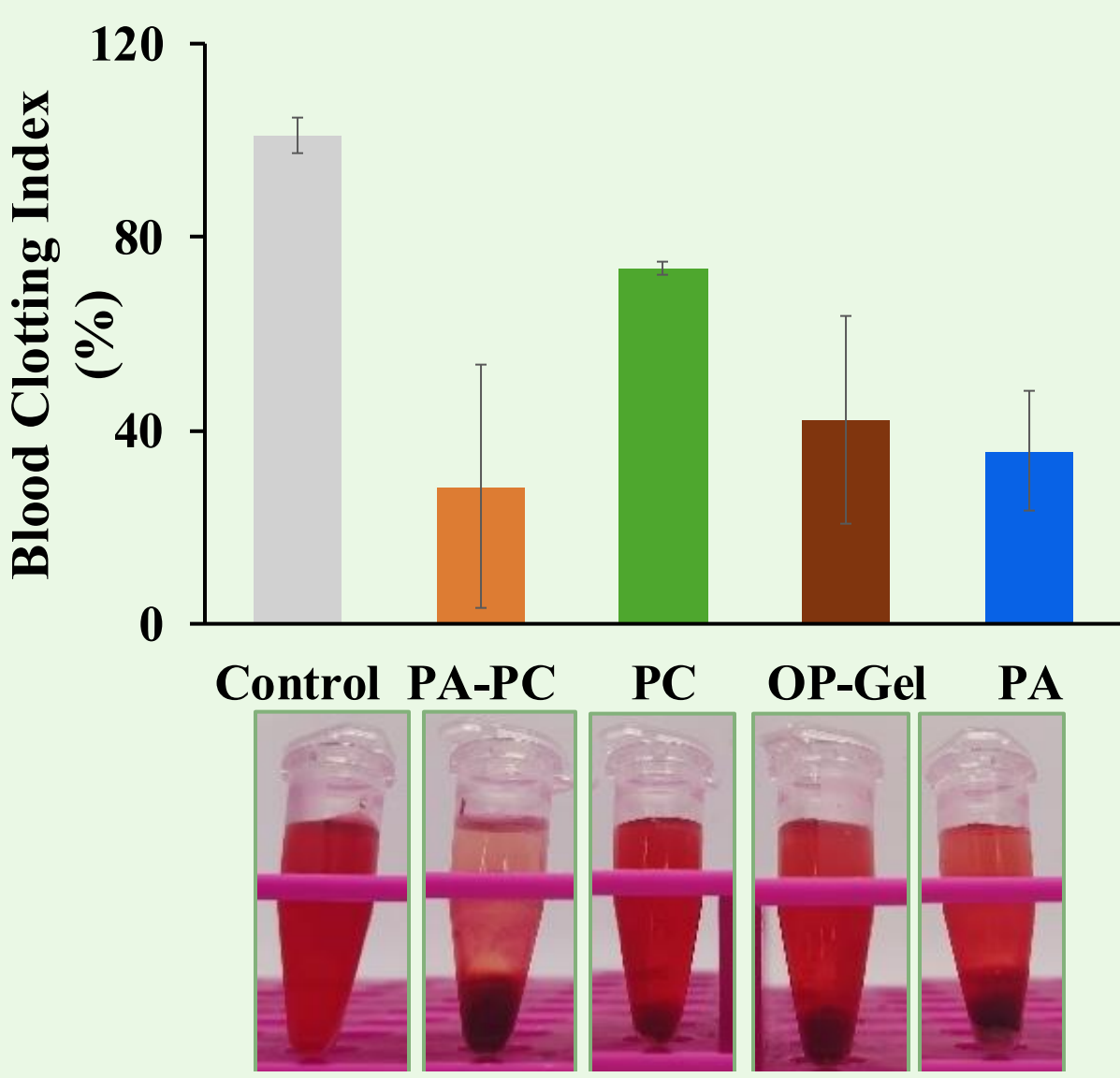


Figure 16. Hemostatic Activity of The Hydrogels

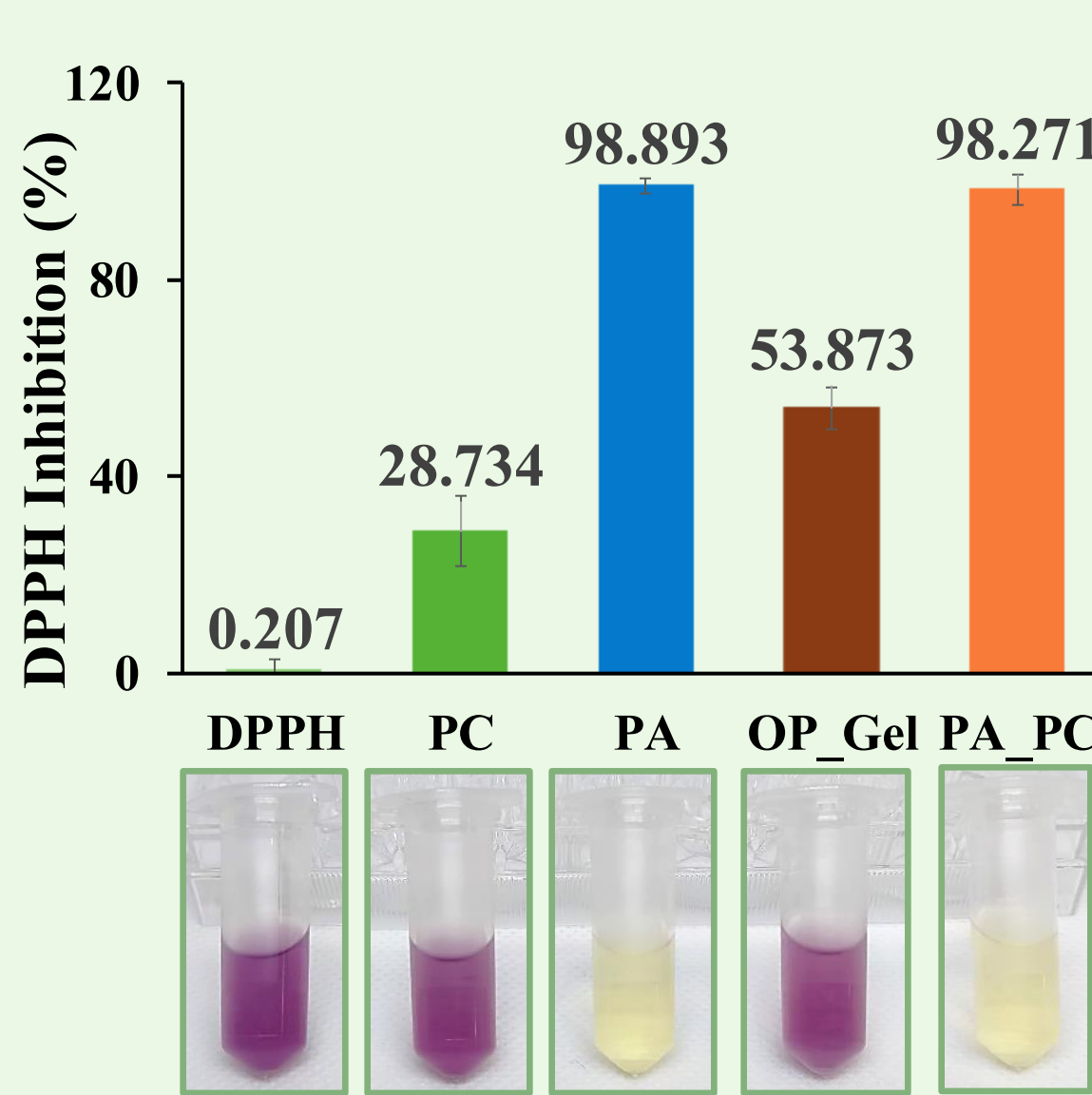


Figure 17. DPPH Analysis



## CONCLUSION

- ✓ FTIR analysis confirmed the oxidation of pectin by the disappearance of the 1588 cm<sup>-1</sup> ester carbonyl peak and the appearance of a new peak at 1606 cm<sup>-1</sup> corresponding to aldehyde C=O stretching [2] .In the hydrogel structure, the formation of new peaks at 1630 cm<sup>-1</sup> (C=N/C=O, amide I) and 1633cm<sup>-1</sup> (N-H bending, amide II) indicated successful Schiff base crosslinking between oxidized pectin and gelatin [3] .
- ✓ The **porous and interconnected microstructure** observed under SEM provides a favorable environment for fluid absorption, nutrient exchange, and diffusion-controlled molecule transport. Micro-scale surface roughness and occasional cracks may contribute to enhanced mechanical interactions and controlled release behavior.
- ✓ Rheological measurements demonstrated high storage modulus (G' > G'') in the linear viscoelastic region, low damping factor (tan δ), and a steep decrease in complex viscosity (n ≈ 0.999) with increasing angular frequency. These results **confirm strong viscoelastic behavior and shear-thinning** properties, enabling injectability and mechanical responsiveness.
- ✓ The hydrogel exhibited efficient **self-healing** behavior due to dynamic Schiff base bonds, and moderate **tissue adhesion** strength based on lap shear testing, indicating structural integrity and potential interface stability in wound environments.
- ✓ Approximately 70% of the encapsulated PC was released in a **controlled manner** within 4 hours. Additionally, **antioxidant capacity** significantly increased in PC and PA-modified formulations, with DPPH inhibition rising from 58.87% (OP\_Gel) to **98.89%** (PA-PC), highlighting its therapeutic potential.
- ✓ Hemolysis results confirmed safe blood compatibility. **Hemostatic activity** improved in PA-PC and PA groups due to fluid uptake and the porous structure that concentrates clotting factors.

## ACKNOWLEDGEMENT

This study supported by ITU-BAP (Scientific Research Projects Unit of Istanbul Technical University) MAB-2024-4551 'Injectable, Self-Healing VEINOGRAP Hydrogel with Pro-Regenerative Activity Toward Vascular Tissue Repair' Project.

## REFERENCES

