

The Activated Platelet-rich Plasma-Fibrin Hydrogel Attenuates Uterine Endometrial Fibrogenesis after Acute Injury via Modulation of Senescence, Inflammation, and Epithelial-Mesenchymal Transition

Nanum Chung^{1,2}, Heeseon Yang^{1,2}, Jungwoo Shin^{1,2}, Hyewon Min^{1,2}, Chungmo Yang^{2,4}, Kangwon Lee⁵, Jung Ryeol Lee^{1,2,3}.

1. Department of Translational Medicine, Seoul National University College of Medicine, Seoul 03080, Republic of Korea

2. Department of Obstetrics and Gynecology, Seoul National University Bundang Hospital, Seongnam 13620, Republic of Korea

3. Department of Obstetrics and Gynecology, Seoul National University College of Medicine, Seoul 03080, Republic of Korea

4. Translational Materials Innovation Group, Institute for Digital Molecular Analytics and Science (IDMxS), Nanyang Technological University, 637553, Singapore

5. Department of Applied Bioengineering, Graduate School of Convergence Science and Technology, Seoul National University, Seoul, 08826, Republic of Korea

In fibrotic diseases, cellular senescence plays a major part by exhibiting healthy or pathologic features, and these characteristics determine whether an organ will maintain its health or progress to disease. Although several studies on fibrotic senescence-related mechanisms have been conducted in organs such as the lungs and liver, research on endometrial fibrotic diseases remains poorly understood. We identified senescence in intrauterine adhesion, a typical fibrotic disease of the uterine endometrium, as fibrosis progressed. Additionally, we applied platelet-rich plasma-fibrin hydrogel (PFH), which demonstrated beneficial effects in our previous study, to prevent fibrosis formation. After inducing endometrial fibrosis through acute exposure to hydrochloric acid, senescence in the endometrium showed a tendency to increase over time along with fibrosis formation. In the experimental group, PFH was infused into the endometrial cavity immediately after damage, and it was confirmed that senescent cells and fibrotic tissues were significantly reduced. These results indicate that when PFH is administered promptly after endometrial damage, pathologic fibrosis is delayed by preventing senescent cell fate, leading to proper wound healing.