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# **Three-Dimensional Printing of rhBMP-2-Loaded LFA Collagen Scaffold with Long Term Delivery for Enhanced Bone Regeneration.**

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Three-Dimensional printing-based scaffolds have demonstrated a remarkable potential for bone formation, regeneration and healing. Research interest in 3D printing-based scaffold is now focused on how to functionalize the scaffolds to accelerate the bone healing process. Recombinant human bone morphogenetic protein-2(rhBMP-2) is recognized as the most potent powerful osteoinductive ability. The rhBMP-2, which is clinically approved by the FDA, KFDA, has been used for various clinical application such as open fracture of mandible, osteomyelitis and osteonecrosis. Despite its efficacy, it has been reported that sustained and local delivery of rhBMP-2 using a suitable carrier is essentially required to accelerated bone healing.

Recently, a variety of sustained rhBMP-2 delivery systems based on 3D printing technology have been introduced. The 3D printing-based collagen scaffold was immersed into rhBMP-2 solution for a delivery system. However, the dipping method could cause inconsistent loading quantity of rhBMP-2 in the scaffold and a waste of rhBMP-2 exceeding the amount necessary. We developed a 3D printing-based rhBMP-2 delivering scaffold only clinically relevant biomaterials and processes using a multi-head deposition system(MHDS).

We have studied Lidocaine-Fibrinogen-Aprotinin(LFA) collagen scaffold loaded rhBMP-2 in 3D printing-based tissue engineering system. Biphasic release of rhBMP-2 could continue for more than 21 days, and keep its osteoinductivity to induce osteogenic differentiation of bone cells. Especially, LFA collagen scaffold system can simultaneously achieve localized long-term controlled release of rhBMP-2 and bone regeneration, which provides a promising route for improving the treatment of bone defects and bone diseases(osteomyelitis, BRONJ, MRONJ, osteoradionecrosis etc.)