# 論 文 要 旨

Enhanced bone formation of calvarial bone defects by lowintensity pulsed ultrasound and recombinant human bone morphogenetic protein-9: a preliminary experimental study in rats

低出力超音波パルスとヒト組換え骨形成タンパク質-9による 頭蓋骨骨欠損部の骨形成促進: ラットを用いた予備実験

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### **Abstract**

# **Objectives**

The aim of this study was to evaluate the combined effects of recombinant human bone morphogenetic protein-9 (rhBMP-9) loaded onto absorbable collagen sponges (ACS) and low-intensity pulsed ultrasound (LIPUS) on bone formation in rat calvarial defects.

#### Materials and Methods

Circular calvarial defects were surgically created in 18 Wistar rats, which were divided into LIPUS-applied (+) and LIPUS-non-applied (-) groups. The 36 defects in each group received ACS implantation (ACS group), ACS with rhBMP-9 (rhBMP-9/ACS group), or surgical control (control group), yielding the following six groups: ACS (+/-), rhBMP-9/ACS (+/-), and control (+/-). The LIPUS-applied groups received daily LIPUS exposure starting immediately after surgery. At 4 weeks, animals were sacrificed and their defects were investigated histologically and by microcomputed tomography.

## Results

Postoperative clinical healing was uneventful at all sites. More new bone was observed in the LIPUS-applied groups compared with the LIPUS-non-applied groups. Newly formed bone area (NBA)/total defect area (TA) in the ACS (+) group ( $46.49\pm7.56\%$ ) was significantly greater than that observed in the ACS (-) ( $34.31\pm5.68\%$ ) and control (-) ( $31.13\pm6.74\%$ ) groups (p<0.05). The rhBMP-9/ACS (+) group exhibited significantly greater bone volume, NBA, and NBA/TA than the rhBMP-9/ACS (-) group ( $2.46\pm0.65$  mm<sup>3</sup> vs.,  $1.76\pm0.44$  mm<sup>3</sup>,  $1.25\pm0.31$  mm<sup>2</sup> vs.,  $0.88\pm0.22$  mm<sup>2</sup>, and  $62.80\pm11.87\%$  vs.,  $42.66\pm7.03\%$ , respectively) (p<0.05). Furthermore, the rhBMP-9/ACS (+) group showed the highest level of bone formation among all groups.

## Conclusion

Within their limits, it can be concluded that LIPUS had osteopromotive potential and enhanced rhBMP-9-induced bone formation in calvarial defects of rats.