

## Effects of a noble metal alloy coating on fibroblast-mediated tissue repair

Aisha S. Ahmed 1, Amina Hamzatova 1, Stefan Grass 2\*, Katrin Ljunggren 2, Vaso Basinou 2, Paul W. Ackermann 1

1. Department of Molecular Medicine and Surgery, Karolinska Institutet, 171 76 Stockholm, Sweden

2. Bactiguard AB, Tullinge, Sweden. \*corresponding author: stefan.grass@bactiguard.com

Disclosures: SG, KL and VB are full time employees of Bactiguard. The study has been funded by Bactiguard.

### OBJECTIVES

- Implant-associated infections are serious complications in orthopedic surgery, with potential severe consequences for peri-implant tissue healing and osseointegration
- A noble metal alloy coating of orthopedic implants has been developed (Bactiguard®, Sweden)
- This coating has been shown to reduce device-related infections by preventing microbial adhesion to the surface
- The objectives of this study were to assess the in vitro effects of Bactiguard-coating on fibroblast proliferation, migration, and collagen production.

### METHODS

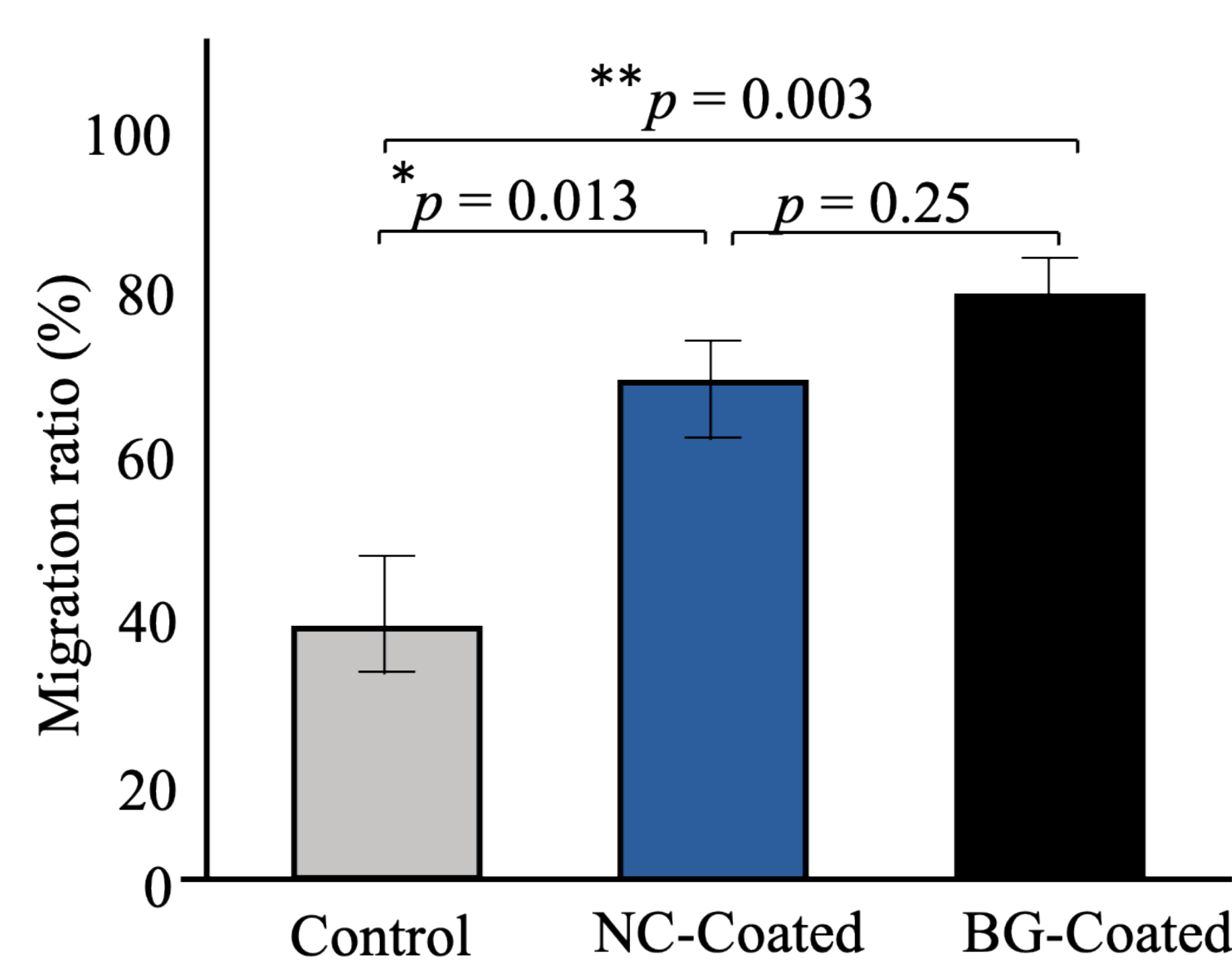
- Human fibroblasts (fHDF/TERT166) were cultured with 1.) Bactiguard-coated titanium coins, 2.) non-coated titanium coins and 3.) without any coins as controls.
- Cell proliferation was studied by counting the number of cells at different timepoints after seeding in the presence of Bactiguard-coated and non-coated titanium coins.
- A cell scratch assay was applied to mimic tissue injury. Cell migration was determined by the rate of scratch recovery in Bactiguard-coated, non-coated titanium coins and control cultures.
- Western blot analysis was used to study Collagen I and III expression in protein lysates extracted from cultures with Bactiguard-coated, non-coated titanium coins and normal controls.
- All statistical analyses were performed using SPSS software (IBM SPSS, v26.0). Between group comparisons were made by the Student's t-test or one-way analysis of variance followed by LSD.  $p < 0.05$  was considered significant.

### RESULTS

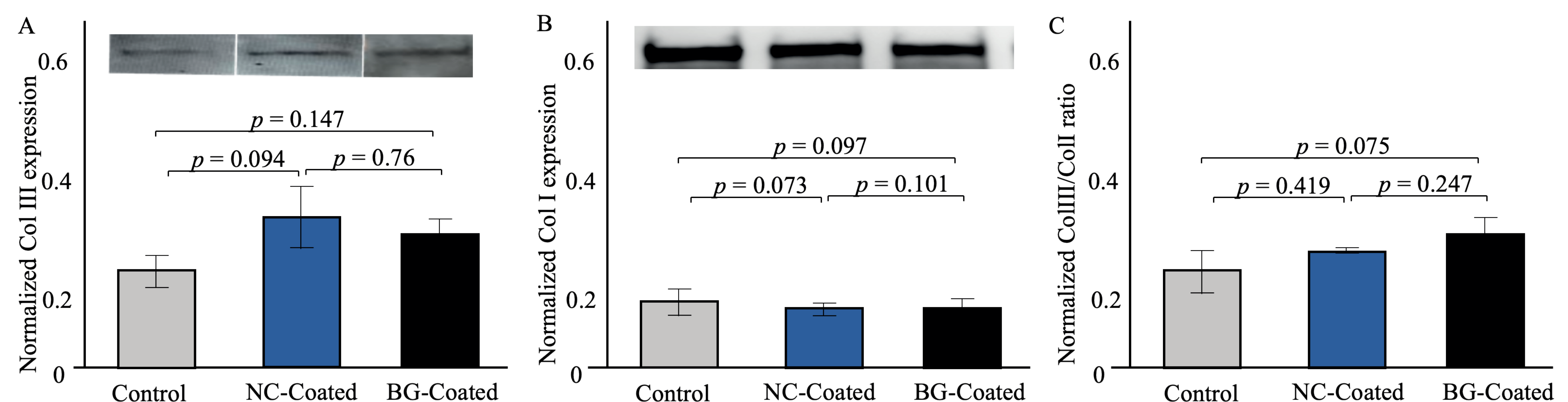
- Cell proliferation experiments at 24 or 36 hours detected no significant differences between the number of cells in cultures with BG-coated and NC titanium coins.
- Significantly higher migration rate was observed in both the Bactiguard-coated (54%,  $p = 0.003$ ) and non-coated coin (46%,  $p = 0.013$ ) cultures compared to the normal control (Fig 1).
- Western blot analysis revealed no significant differences among Bactiguard-coated, non-coated coins or control cultures in the amount of Collagen I and Collagen III expression (Fig 2A, B)

### CONCLUSIONS

- The study results of this in vitro model on fibroblast-mediated tissue repair suggest that the Bactiguard-coating does not exert any adverse effects on tissue repair.
- The higher migration ratio and slightly higher Collagen III/Collagen I ratio on the coin cultures (i.e., Bactiguard coated and non-coated) compared to normal control, indicates inflammatory healing response from placing a coin on a fibroblast cell culture.
- Coated implants can help reduce infections but may increase the risk of toxicological effects. Thus, non-toxic, infection-prevention coatings that do not disrupt the normal cell repair mechanisms are preferable.



**Fig 1.** Semi-quantitative analysis of cell migration ratio in control, Non-Coated (NC) and Bactiguard (BG)-coated cultures at 24 hrs. Data reported as mean  $\pm$  SEM with  $n = 3$  replicates. \* $p < 0.05$ , \*\* $p < 0.005$ .



**Fig 2.** The representative western blot images and semi-quantitative analysis of A) Col III, B) Col I, and C) Col III/I ratio from protein lysates generated by control, Non-Coated (NC) and Bactiguard (BG)-coated cultures at 36 hrs. Signal intensity was used for analysis and the intensity of the house-keeping gene (beta-actin) used for normalization. Data reported as mean  $\pm$  SEM with  $n = 3$  replicates.



### References:

1. Karupiah, Thevarajan, Aik Peng Yong, Ze Wee Ong, Heng Keat Tan, Wei Chern Tang, and Hishamuddin Bin Salam. 2022. Use of a Novel Anti-Infective Noble Metal Alloy-Coated Titanium Orthopedic Nail in Patients with Open Fractures: A Case Series from Malaysia. *Antibiotics* 11, no. 12: 1763.
2. Suska, F.; Svensson, S.; Johansson, A.; Emanuelsson, L.; Karlholm, H.; Ohrlander, M.; Thomsen, P. 2010. In vivo evaluation of noble metal coatings. *J. Biomed. Mater. Res. B Appl. Biomater.* 92, 86–94.
3. Chen, J., Wang, J., Hart, D. A., Ahmed, A. S., and Ackermann, P. W. 2022 Complement factor D as a predictor of Achilles tendon healing and long-term patient outcomes. *FASEB J* 36, e22365.
4. Zou SB, Yoon WY, Han SK, Jeong SH, Cui ZJ, Kim WK. Cytotoxicity of silver dressings on diabetic fibroblasts. *Int Wound J*. 2013;10(3):306-312.