



## Surface modifications by gas plasma control osteogenic differentiation of MC3T3-E1 cells

Ana M.C. Barradas<sup>a</sup>, Kristina Lachmann<sup>b</sup>, Gregor Hlawacek<sup>c</sup>, Cathelijne Frielink<sup>d</sup>, Roman Truckenmoller<sup>a</sup>, Otto C. Boerman<sup>d</sup>, Raoul van Gastel<sup>c</sup>, Henk Garritsen<sup>e,f</sup>, Michael Thomas<sup>b</sup>, Lorenzo Moroni<sup>a</sup>, Clemens van Blitterswijk<sup>a</sup>, Jan de Boer<sup>a,\*</sup>

<sup>a</sup> Department of Tissue Regeneration, MIRA Institute for Biomedical Research and Technical Medicine, University of Twente, PO Box 217, 7500 AE Enschede, The Netherlands

<sup>b</sup> Fraunhofer Institute for Surface Engineering and Thin Films IST, Bienroder Weg 54 E, 38108 Braunschweig, Germany

<sup>c</sup> Physics of Interfaces and Nanomaterials, MESA+ Institute for Nanotechnology, University of Twente, PO Box 217, 7500 AE Enschede, The Netherlands

<sup>d</sup> Department of Nuclear Medicine, Radboud University Nijmegen Medical Centre, Geert Grooteplein 8, 6525 GA Nijmegen, The Netherlands

<sup>e</sup> Helmholtz Centre for Infection Research GmbH, Inhoffenstraße 7, 38124 Braunschweig, Germany

<sup>f</sup> Department of Transfusion Medicine, Städtisches Klinikum, Celler Straße 38, 38114 Braunschweig, Germany

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### ABSTRACT

Numerous studies have shown that the physicochemical properties of biomaterials can control cell activity. Cell adhesion, proliferation, differentiation as well as tissue formation in vivo can be tuned by properties such as the porosity, surface micro- and nanoscale topography and chemical composition of biomaterials. This concept is very appealing for tissue engineering since instructive properties in bioactive materials can be more economical and time efficient than traditional strategies of cell pre-differentiation in vitro prior to implantation. The biomaterial surface, which is easy to modify due to its accessibility, may provide the necessary signals to elicit a certain cellular behavior. Here, we used gas plasma technology at atmospheric pressure to modify the physicochemical properties of polylactic acid and analyzed how this influenced pre-osteoblast proliferation and differentiation. Tetramethylsilane and 3-aminopropyl-trimethoxysilane with helium as a carrier gas or a mixture of nitrogen and hydrogen were discharged to polylactic acid discs to create different surface chemical compositions, hydrophobicity and microscale topographies. Such modifications influenced protein adsorption and pre-osteoblast cell adhesion, proliferation and osteogenic differentiation. Furthermore polylactic acid treated with tetramethylsilane enhanced osteogenic differentiation compared to the other surfaces. This promising surface modification could be further explored for potential development of bone graft substitutes.

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### 1. Introduction

Bone tissue engineering has emerged as a field providing alternatives to autologous bone grafts, which are still considered the gold standard treatment for healing bone defects [1–3]. Tissue-engineering strategies focus on the development of scaffolds and/or on the combination of scaffolds with cells. Traditionally cells are pre-differentiated into an osteogenic lineage through addition of growth factors or steroids, such as bone morphogenetic proteins (BMPs) or dexamethasone [4–8]. Alternatively, cell differentiation may be controlled by the physicochemical properties of the scaffold material [9–11]. This represents a more economic and expeditious approach and has the additional advantage that biologically

relevant molecular signals are still transmitted to the cells through cell–surface interactions after the graft has been implanted. For example, induction of bone formation is known to be influenced by the pore size of biphasic calcium phosphate ceramic granules [12], the depth of surface concavities in hydroxyapatite ceramic discs [13] and the chemical composition of the ceramic materials [14]. This demonstrates the relevance of material properties for clinical application.

In addition to changing bulk properties of a biomaterial, one can also change the surface properties, such as topography or chemistry. For instance, it has been noted that NH<sub>2</sub>-enriched surfaces promoted osteogenesis of human bone marrow derived mesenchymal stromal cells (hMSCs), whereas chondrogenesis was favored by COOH and OH groups [15]. However, Phillips et al. could not ascribe expression of chondrogenic markers to one specific group [16]. Changes in surface chemistry are accompanied by differences in material–protein interaction, which may account for the observed

\* Corresponding author. Tel.: +31 53 489 3400; fax: +31 53 489 2150.

E-mail address: [j.deboer@utwente.nl](mailto:j.deboer@utwente.nl) (J. de Boer).