## Engineering Surface Chemistries to Improve Tissue-Material Integration in Medical Implants

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Tissue-material integration is a key challenge of engineering medical implants and implantable biosensors. Implants often undergo an intense foreign body reaction orchestrated by human macrophages (hM $\phi$ s), leading to severe inflammation and expensive revision surgeries, potentially interfering with the functioning of an implanted biosensor. This in vitro study investigated the potential of low-cost porous silicone-gel scaffolds with various mussel-inspired polydopamine (PDA)-based surface coatings as medical implant surfaces. Scaffolds of varying pore sizes (50-500µm) were modified with PDA, cell adhesive RGD peptide, and growth factor TGF- $\beta$ . The proliferation and extracellular matrix (ECM) secretion of human dermal fibroblasts (hDFs) on these scaffolds were studied in the presence and absence of hM $\phi$ s, yielding a relatively holistic in vitro model of the tissue regeneration process around the implant. Three separate investigations were conducted. The first measured cytokine release profiles of hM $\phi$ s cultured on scaffolds, showing that PDA-based coatings halved release of TNF- $\alpha$  (a major pro-inflammatory cytokine). The second experiment refined and validated the novel "flipped-floater" system for co-culturing hDFs and hM $\phi$ s. The third experiment studied hDFs both alone and co-cultured with hM $\phi$ s using the flipped-floater system. hDFs on their own exhibited around twice the proliferation on PDA as on uncoated scaffolds, and PDA provided a seven-fold boost with hM $\phi$ s present. 100-300µm pores yielded the best performance. SEM pictures show hDFs growing up to 500µm deep and secreting healthy ECM on PDA-coated scaffolds. These results indicate that PDA coatings with properly-structured scaffolds have significant potential to improve tissue-material integration in medical implants.