

Biomaterial Surface Functionalization For Enhanced Cell Adhesion

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Introduction

The surface functionalization of biomaterials stands as a cornerstone in advancing tissue engineering and regenerative medicine, primarily by enhancing cellular adhesion, a critical initial step for successful integration and function of engineered tissues. This process involves the strategic modification of biomaterial surfaces with specific molecules, such as peptides, proteins, or components of the extracellular matrix (ECM), to cultivate an environment conducive to cell attachment, proliferation, and differentiation. The methodologies employed span a spectrum from physical techniques like plasma treatment and grafting to sophisticated chemical modifications and the incorporation of bioactive factors, all aimed at replicating the natural cellular milieu to guide cellular behavior and ultimately improve the efficacy of engineered constructs [1].

A principal objective in the design of biomaterials for cell adhesion is the accurate mimicry of the extracellular matrix (ECM). Research in this area focuses on precisely controlling surface topography and chemistry, utilizing advanced techniques such as photolithography and controlled surface grafting, to meticulously dictate cellular responses. The findings highlight how nanoscale features and the specific presentation of RGD motifs on material surfaces can profoundly enhance mesenchymal stem cell adhesion and subsequent differentiation into osteogenic lineages, thereby underscoring the critical importance of bioinspired surface design for regenerative applications [2].

The strategic presentation of specific cell-binding ligands, particularly peptides derived from ECM proteins, represents a highly effective approach for promoting cell adhesion to biomaterials. Investigations into this area have explored the efficacy of integrating RGD sequences into hydrogel scaffolds, demonstrating that the density and spatial arrangement of these peptides significantly influence fibroblast adhesion, spreading, and proliferation. This body of work offers valuable insights into optimizing peptide-based surface functionalization strategies for enhanced biocompatibility and improved tissue integration [3].

Plasma treatment has emerged as a versatile and effective technique for the surface modification of biomaterials. Studies employing atmospheric pressure plasma have successfully functionalized polymer surfaces, introducing polar groups that promote protein adsorption and, consequently, enhance cell adhesion. Empirical evidence shows that specific plasma parameters can be manipulated to control surface chemistry, thereby optimizing the attachment of endothelial cells, a factor of paramount importance in vascular tissue engineering. This method presents an attractive solvent-free alternative for enhancing biomaterial-cell interactions [4].

The development of bioactive coatings capable of actively promoting cell adhesion and integration is paramount for the successful deployment of implantable devices.

Recent advancements include the synthesis and characterization of novel coatings that incorporate heparin and growth factors onto titanium implant surfaces. The results from such studies consistently demonstrate a significant improvement in osteoblast adhesion, proliferation, and differentiation when compared to uncoated surfaces, indicating that bio-inspired approaches hold considerable promise for enhancing osseointegration and overall implant success [5].

Hydrogel biomaterials are extensively utilized in tissue engineering owing to their inherent biocompatibility and their remarkable ability to recapitulate the native ECM. However, surface modification of these hydrogels is essential for precisely controlling cellular interactions. Current research is focused on developing photocrosslinkable hydrogel systems that can be readily functionalized with cell-adhesive peptides, such as RGD, and growth factors using light-activated processes. These studies have shown that the controlled presentation of such cues on the hydrogel surface effectively promotes robust stem cell adhesion and osteogenic differentiation [6].

The interplay between cells and implantable materials is profoundly influenced by the surface properties of the materials themselves. Exploration into versatile platforms for surface functionalization has led to the development of mussel-inspired polydopamine (PDA) coatings. These PDA coatings are readily modifiable with a diverse array of biomolecules, leading to enhanced adhesion and proliferation of human dermal fibroblasts. The efficacy of PDA coatings in effectively immobilizing collagen, thereby improving cell attachment, highlights their significant potential for applications in wound healing [7].

Chitosan-based biomaterials, while promising for tissue engineering applications, frequently necessitate surface modification to improve their inherent cell adhesion capabilities. Investigations into this area have explored the use of simple glutaraldehyde crosslinking methods to immobilize fibronectin onto chitosan films. The resulting fibronectin-coated films exhibit substantially enhanced adhesion and spreading of pre-osteoblast cells when contrasted with unmodified chitosan, offering a straightforward strategy for augmenting the bioactivity of chitosan scaffolds for bone regeneration [8].

Precision in controlling the presentation of biomolecular cues on material surfaces is fundamental to effectively directing cellular behavior. This research has leveraged click chemistry to achieve the precise anchoring of peptides containing cell-adhesion motifs onto polymer surfaces. The findings demonstrate that the density of these anchored peptides can be meticulously tuned, enabling controlled cell adhesion and subsequent differentiation of mesenchymal stem cells towards a neuronal lineage. This sophisticated approach offers a high degree of control over the cell-instructive microenvironment [9].

Nanoparticle-based surface functionalization presents unique advantages for augmenting cell-biomaterial interactions. This study highlights the utilization of gold

nanoparticles (AuNPs) functionalized with cell-adhesive peptides to coat biomaterial surfaces. The resulting AuNP coating significantly enhances the adhesion and spreading of human umbilical vein endothelial cells (HUVECs), underscoring the potential of nanoparticle-mediated surface modification for improving the performance of both biomedical devices and tissue engineering scaffolds [10].

Description

Surface engineering of biomaterials is paramount for enhancing cell adhesion, a crucial step in tissue engineering and regenerative medicine. This involves modifying biomaterial surfaces with specific molecules like peptides, proteins, or extracellular matrix (ECM) components to create a favorable microenvironment that promotes cell attachment, proliferation, and differentiation. Strategies range from physical methods like plasma treatment and grafting to chemical modifications and the incorporation of bioactive factors, aiming to mimic the natural cellular environment and guide cellular behavior [1].

Mimicking the extracellular matrix (ECM) is a primary objective in biomaterial design for cell adhesion. This involves precisely controlling surface topography and chemistry using techniques such as photolithography and controlled surface grafting to dictate cell behavior. Specifically, nanoscale features and the presentation of RGD motifs can significantly enhance mesenchymal stem cell adhesion and subsequent differentiation into osteogenic lineages, emphasizing the importance of bioinspired surface design for regenerative applications [2].

The presentation of specific cell-binding ligands, such as peptides derived from ECM proteins, is a powerful method for promoting cell adhesion to biomaterials. Research has investigated the efficacy of incorporating RGD sequences into hydrogel scaffolds, demonstrating that peptide density and spatial arrangement significantly influence fibroblast adhesion, spreading, and proliferation. This work offers insights into optimizing peptide-based surface functionalization for improved biocompatibility and tissue integration [3].

Plasma treatment serves as an effective and versatile technique for the surface modification of biomaterials. Atmospheric pressure plasma has been utilized to functionalize polymer surfaces, introducing polar groups that enhance protein adsorption and subsequent cell adhesion. Specific plasma parameters can control surface chemistry, optimizing endothelial cell attachment, which is critical for vascular tissue engineering. This method provides a solvent-free approach for improving biomaterial-cell interactions [4].

The development of bioactive coatings that actively promote cell adhesion and integration is essential for implantable devices. Novel coatings incorporating heparin and growth factors onto titanium implant surfaces have been synthesized and characterized. These functionalized surfaces significantly improve osteoblast adhesion, proliferation, and differentiation compared to uncoated surfaces, suggesting that bio-inspired approaches hold promise for enhancing osseointegration and implant success [5].

Hydrogel biomaterials are widely employed in tissue engineering due to their biocompatibility and ability to mimic the native ECM. Surface modification of these hydrogels is critical for controlling cell interactions. Studies focus on developing photocrosslinkable hydrogel systems that can be functionalized with cell-adhesive peptides (like RGD) and growth factors using light. These methods demonstrate that controlled surface presentation of these cues promotes robust stem cell adhesion and osteogenic differentiation [6].

The interaction of cells with implantable materials is heavily influenced by surface properties. Mussel-inspired polydopamine (PDA) coatings offer a versatile platform for surface functionalization. PDA coatings can be easily modified with

various biomolecules, promoting enhanced adhesion and proliferation of human dermal fibroblasts. The study demonstrates PDA coatings' ability to effectively immobilize collagen, leading to improved cell attachment and showcasing their potential for wound healing applications [7].

Chitosan-based biomaterials require surface modification to improve cell adhesion. This work investigated the use of glutaraldehyde crosslinking to immobilize fibronectin onto chitosan films. The fibronectin-coated films showed significantly enhanced adhesion and spreading of pre-osteoblast cells compared to unmodified chitosan, presenting a straightforward strategy to improve the bioactivity of chitosan scaffolds for bone regeneration [8].

Controlling the presentation of biomolecular cues on material surfaces is key to directing cell behavior. Click chemistry has been employed to precisely anchor peptides containing cell-adhesion motifs onto polymer surfaces. The density of these peptides can be finely tuned, leading to controlled cell adhesion and subsequent differentiation of mesenchymal stem cells towards a neuronal lineage, offering precise control over the cell-instructive microenvironment [9].

Nanoparticle-based surface functionalization offers unique advantages for enhancing cell-biomaterial interactions. Gold nanoparticles (AuNPs) functionalized with cell-adhesive peptides were used to coat a biomaterial surface. The AuNP coating promoted significantly enhanced adhesion and spreading of human umbilical vein endothelial cells (HUVECs), highlighting the potential of nanoparticle-mediated surface modification for improving the performance of biomedical devices and tissue engineering scaffolds [10].

Conclusion

Surface functionalization of biomaterials is essential for improving cell adhesion, a critical process in tissue engineering and regenerative medicine. This involves modifying material surfaces with molecules like peptides and proteins to encourage cell attachment, proliferation, and differentiation. Various techniques are employed, including physical and chemical modifications, as well as the incorporation of bioactive factors. Research focuses on mimicking the extracellular matrix through precise control of surface topography and chemistry, utilizing methods like photolithography and peptide immobilization (e.g., RGD sequences). Plasma treatment and bioactive coatings are also effective for enhancing cell adhesion to implantable devices. Hydrogels and chitosan-based materials are modified using techniques such as photocrosslinking and fibronectin immobilization. Nanoparticle-based approaches, particularly using gold nanoparticles, show promise for improving endothelial cell adhesion. Overall, these strategies aim to create more biocompatible and cell-instructive biomaterial surfaces for improved therapeutic outcomes.

Acknowledgement

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Conflict of Interest

None.

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