

# Regenerative medicine and osteoarthritis: SVF, exosomes and beyond

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## **Torbjörn Ogéus from Stockholms Led- & Smärtspecialist guides us through the evolving science of stromal vascular fraction, exosomes, and beyond in regenerative medicine and osteoarthritis**

Osteoarthritis (OA), a chronic degenerative joint condition, affects hundreds of millions worldwide and remains a leading cause of disability and reduced quality of life. While traditional treatments focus on symptom control – primarily pain management and surgical intervention – the field of regenerative medicine is rapidly rewriting the narrative.

Recent advancements in cellular therapy, particularly stromal vascular fraction (SVF), exosomes, and novel biomaterials, provide hope for true biological repair. This article delves into the cutting-edge developments in this field, highlighting new findings, the promise of alternative therapies, and the roadblocks posed by outdated regulatory frameworks.

### **SVF and exosomes: Changing the landscape of OA treatment**

SVF, a heterogeneous mixture of cells derived from adipose tissue, has emerged as a powerful regenerative agent. Rich in mesenchymal stem cells, pericytes, endothelial progenitor cells, and immune-modulatory factors, SVF can stimulate cartilage repair, reduce inflammation, and improve joint function.

In the retrospective study “Stromal Vascular Fraction with Platelet-Rich Fibrin for Osteoarthritis Management in Knee and Hip Osteoarthritis: A Retrospective 2-Year Follow-Up Study,” we described the synergistic use of SVF and platelet-rich fibrin (PRF) which demonstrated significant clinical improvements over a sustained period.

This combination not only harnessed the regenerative capacity of stem cells but also leveraged PRF’s ability to provide a fibrin scaffold, ensuring cell retention and a prolonged release of growth factors. Patients experienced reduced pain and increased mobility, pointing to a potentially disease-modifying approach rather than simple symptom alleviation. <sup>(1)</sup>

More recently, exosomes – nanoscale extracellular vesicles released by stem cells – have gained traction as a therapeutic frontier. Unlike their parent stem cells, exosomes carry no risk of differentiation-related complications and exert potent paracrine effects. Earlier this year, we published the study “Amniotic Derived Exosomes with Platelet Rich Fibrin Combined in the Treatment of Hip and Knee Osteoarthritis: A Retrospective 1-Year Follow-Up Study”. It provided early but compelling evidence that these tiny vesicles, when

combined with PRF, can yield clinically meaningful outcomes similar to the results of SVF combined with PRF in the abovementioned study. The amniotic origin of the exosomes adds an additional layer of immune privilege and potency, offering a viable option for patients who may not be ideal candidates for autologous cell therapy. <sup>(2)</sup>

### **Safety concerns: The autologous paradox**

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Despite the promising results with autologous therapies like SVF, caution is warranted. The commonly held belief that using one's own cells automatically equates to safety is being increasingly challenged. In the study "Multiple intra-articular injections with adipose-derived stem cells for knee osteoarthritis cause severe arthritis with anti-histone H2B antibody production," repeated intra-articular injections of autologous adipose-derived stem cells resulted in the unexpected development of severe arthritis and an autoimmune-like response with catastrophic results in 50% of the cases. The presence of anti-histone H2B antibodies suggests that autologous cells, when reintroduced multiple times, may provoke an adverse immune reaction, especially when inflammatory joint environments alter cellular surface markers.

This raises critical questions about the one-size-fits-all regulatory mandates in regions like the European Union (EU). Current EU legislation dictates that all stem cell therapies for OA must be autologous. While well-intentioned to prevent transmission of infections and immune complications from donor cells, this requirement overlooks the nuances of immunogenicity and biological tolerance. In reality, allogeneic stem cell products – especially those pooled from multiple donors – can be carefully screened, purified, and standardised, offering a lower risk of immunogenicity and batch variability. The blanket restriction against allogeneic stem cells stifles innovation and may paradoxically increase patient risk in certain scenarios. <sup>(3)</sup>

### **Emerging innovations: Hydrogels and blood-derived stem cells**

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Alongside these cellular advances, the field is witnessing rapid developments in delivery systems and support matrices. Hydrogels, with their viscoelastic and biocompatible properties, are now being explored as scaffolds and carriers for stem cells and growth factors. These hydrogels mimic the native extracellular matrix, enhancing cell viability, directing differentiation, and maintaining the structural integrity of joint spaces.

Recent studies with thermoresponsive hydrogels, injectable yet solidifying at body temperature, have shown great promise. These carriers enable localised and sustained cell delivery, reduce mechanical washout, and provide a 3D environment conducive to cartilage regeneration. Some are even being functionalised with bioactive molecules to further enhance cellular interaction and signaling pathways. <sup>(4)</sup>

Another promising frontier lies in using a newly discovered subtype of blood stem cells called small blood-derived stem cells (SBSCs), an emerging class of progenitor cells found in peripheral blood. Isolated from autologous blood samples, these cells exhibit

pluripotent properties similar to embryonic stem cells, with the potential to differentiate into multiple musculoskeletal lineages.

Unlike adipose- or bone marrow-derived stem cells, SBSCs are easy to obtain with minimal invasiveness, making them a potentially transformative option for same-day, point-of-care regenerative therapy. Early preclinical data suggests that SBSCs could offer similar efficacy with fewer procedural complications and less regulatory red tape, given their autologous and minimally manipulated nature, without the risk of developing tumours at a fraction of the cost of today's stem cell therapies.

## **Toward a new paradigm in OA therapy**

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Taken together, the growing body of evidence underscores a paradigm shift in the treatment of OA. Rather than being viewed as a chronic, irreversible condition, OA is increasingly seen as a modifiable disease amenable to biologically active interventions.

The combined use of SVF, exosomes, and PRF offers a multifaceted regenerative strategy: reducing inflammation, promoting tissue repair, and restoring joint function.

Yet progress is not purely scientific – it must also be regulatory and philosophical. As the case of autologous stem cell complications demonstrates, safety must be assessed through the lens of real-world outcomes, not theoretical ideals. It is time for regulators to embrace a risk-benefit model grounded in evolving science, allowing carefully screened allogeneic therapies a fair pathway to clinical use.

Simultaneously, adjunct innovations like hydrogels and blood-derived stem cells are expanding the toolkit available to orthopaedic practitioners. These developments suggest that the future of joint regeneration will be increasingly personalised, minimally invasive, and biologically informed.

## **OA: Conclusion**

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The road to a cure for OA is still being paved, but the progress made in regenerative medicine over the past decade is nothing short of remarkable. The studies presented in this article highlight the real-world applicability of cell-based and exosome therapies. However, we must also heed the lessons from immune reactions and regulatory limitations that stand in the way of innovation.

Looking ahead, integrating smart carriers like hydrogels and the rise of SBSCs from peripheral blood may revolutionise the accessibility and efficacy of regenerative therapies. With continued research, open-minded regulation, and interdisciplinary collaboration, we are poised to redefine how osteoarthritis is treated – moving from symptom management to actual biological repair.

1. T. Ogéus. Stromal Vascular Fraction with Platelet-Rich Fibrin for Osteoarthritis Management in Knee and Hip Osteoarthritis: A Retrospective 2-Year Follow-Up Study. *Journal of Orthopedics and Sports Medicine*. 6 (2024): 135-143.

2. T. Ogéus. Amniotic Derived Exosomes with Platelet Rich Fibrin Combined in the Treatment of Hip and Knee Osteoarthritis: A Retrospective 1-Year Follow-Up Study. Journal of Orthopedics and Sports Medicine. 7 (2025): 129-137.
3. Y. Hosono, Y et al. Multiple intra-articular injections with adipose-derived stem cells for knee osteoarthritis cause severe arthritis with anti-histone H2B antibody production. Regenerative therapy, 24 (2023): 147–153.
4. T. Ogéus. Hydrogel alone or in Combination with Regenerative Interventions for Knee Osteoarthritis, A Case Series. Journal of Orthopedics and Sports Medicine. 7 (2025): 1

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