

Fig. 8

**Bone marrow mesenchymal stem cell-derived exosomes protect cartilage damage and relieve knee osteoarthritis pain in a rat model of osteoarthritis**

**SPRINGER NATURE**

**Author:** Lei He et al

**Publication:** Stem Cell Research & Therapy

**Publisher:** Springer Nature

**Date:** Jul 10, 2020

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Title of targeted journal	Bioactive Materials
Publisher	Elsevier
Expected publication date	Oct 2022
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Fig.10

**Exosomes from antler stem cells alleviate mesenchymal stem cell senescence and osteoarthritis**

**SPRINGER NATURE**

**Author:** Jinghui Lei et al  
**Publication:** Protein & Cell  
**Publisher:** Springer Nature  
**Date:** Aug 3, 2021  
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Fig.12

**Curcumin-primed human BMSC-derived extracellular vesicles reverse IL-1 $\beta$ -induced catabolic responses of OA chondrocytes by upregulating miR-126-3p**

**SPRINGER NATURE**

**Author:** Shushan Li et al

**Publication:** Stem Cell Research & Therapy

**Publisher:** Springer Nature

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Fig.13



RESEARCH ARTICLE | Open Access |

## The umbilical cord mesenchymal stem cell-derived exosomal lncRNA H19 improves osteochondral activity through miR-29b-3p/FoxO3 axis

Litao Yan, Gejun Liu, Xing Wu

First published: 13 January 2021 | <https://doi.org/10.1002/ctm2.255> | Citations: 17

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Litao Yan and Gejun Liu contributed equally to this work.

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### Abstract

#### Background

Our previous study revealed that the exosomal lncRNA H19 derived from umbilical cord mesenchymal stem cells (UMSCs) plays a pivotal role in osteochondral regeneration. In this study, we investigated whether the exosomal lncRNA H19 could act as a competing endogenous RNA (ceRNA) to potentiate osteochondral activity in chondrocytes.

#### Methods

Dual-luciferase reporter assay, RNA pull-down, RNA immunoprecipitation (RIP), and fluorescence in situ hybridization (FISH) were carried to verify the interaction between miR-29b-3p and both lncRNA H19 and the target mRNA FoxO3. Chondrocytes were treated with UMSC-derived exosomes, which highly expressing lncRNA H19 expression, followed by apoptosis, migration, senescence, and matrix secretion assessments. An in vivo SD rat cartilage defect model was carried out to explore the role and mechanism of lncRNA H19/miR-29b-3p.



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Title of targeted journal	Bioactive Materials
Publisher	Elsevier
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Fig.14-Right

## Reversing the surface charge of MSC-derived small extracellular vesicles by $\epsilon$ PL-PEG-DSPE for enhanced osteoarthritis treatment

Kai Feng, Xuetao Xie, Ji Yuan, Liangzhi Gong, Zhaochen Zhu, Juntao Zhang, Haiyan Li, Yunlong Yang, Yang Wang

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### Abstract

Mesenchymal stem cell-derived small extracellular vesicles (MSC-sEVs) possess a great therapeutical potential for osteoarthritis (OA) treatment. However, the steric and electrostatic hindrance of cartilage matrix leads to very limited distribution of MSC-sEVs in cartilage and low bioavailability of MSC-sEVs after intra-articular injection. To overcome this, a strategy to reverse the surface charge of MSC-sEVs by modifying the MSC-sEVs with a novel cationic amphiphilic macromolecule namely  $\epsilon$ -polylysine-polyethylene-distearyl phosphatidylethanolamine (PPD) was developed in this study. Through incubation with 100  $\mu\text{g}/\text{ml}$  PPD, positively charged MSC-sEVs (PPD-sEVs) were obtained, and the modification process showed nearly no disturbance to the integrity and contents of sEVs and exhibited good stability under the interference of anionic macromolecules. A more effective cellular uptake and homeostasis modulation ability of PPD-sEVs than unmodified MSC-sEVs to chondrocytes was demonstrated. More importantly, PPD-sEVs demonstrated significantly enhanced cartilage uptake, cartilage penetration, and joint retention capacity as compared to MSC-sEVs. Intra-articular injection of PPD-sEVs into a mouse OA model showed significantly improved bioavailability than MSC-sEVs, which resulted in enhanced therapeutic efficacy with reduced injection frequency. In general, this study provides a facile and effective strategy to improve the intra-articular bioavailability of MSC-sEVs and has a great potential to accelerate the clinical practice of MSC-sEVs based OA therapy.



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### Research Funding



Research Paper

## Chondrocyte-specific genomic editing enabled by hybrid exosomes for osteoarthritis treatment

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