

## **Supplemental Materials**

### **Anti-inflammatory Therapy for Temporomandibular Joint Osteoarthritis Using mRNA Medicine Encoding Interleukin-1 Receptor Antagonist**

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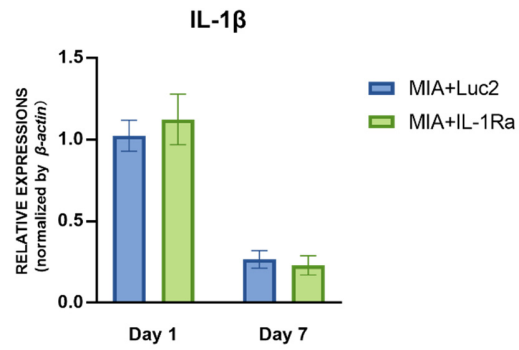
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### Supplemental Figure S1



**Figure S1.** The expression levels of Interleukin-1 $\beta$  (IL-1 $\beta$ ) on the 1<sup>st</sup> day and 7<sup>th</sup> day after the mRNA treatment.  $\beta$ -actin expression was used for normalization. Animal number = 6/group.

**Table S1.** Comparison of physicochemical properties between lipid nanoparticles and polyplex nanomicelles for loading mRNA.

	<b>Lipid Nanoparticles-mRNA</b>	<b>Polyplex Nanomicelles</b>
<b>Components</b>	Cationic/ionizable lipids and helper lipids such as phospholipids, cholesterol, or polyethylene glycol (PEG)-functionalized lipids (PEG-lipids) [15, 16]	Block copolymers composed of PEG and cationic polyamino acid [3-5]
<b>Structure</b>	A core-shell structure with mRNA encapsulated in the core in general but highly variable in structure [19-20]	A core-shell structure surrounded by dense PEG palisade with mRNA encapsulated in the core [3-5]
<b>Assembling</b>	Self-assembled mainly driven by hydrophobic interactions and guided by electrostatic interactions [18]	Self-assembled by electrostatic interactions [3-5]
<b>Size</b>	Changeable based on applications and components [18, 21]	Regulated between 50-100 nm [6]
<b>Administration route</b>	Applicable for various routes e.g., intravenous (i.v.) and local administration.	Various Local administration routes [9-15] and hydrodynamic intravenous injection [8]
<b>Biodistribution after i.v. administration</b>	Accumulations in the liver and lymph nodes [22, 23]	-
<b>Immune response after administration</b>	Considerable immune responses, which could work as adjuvant for vaccination [1, 2]	Less or no immune response [6, 7, 9, 14]

#### References for Table S1

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