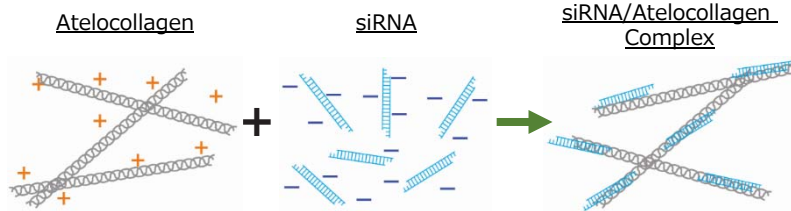


Local & Systemic Use: Simple, Efficient Method for Transfection of siRNA and miRNA with low toxicity

Outline

Atelocollagen, the main component of AteloGene®, forms siRNA/miRNA atelocollagen complexes by appropriate mixing with synthetic siRNA/miRNA. The siRNA/miRNA atelocollagen complexes are optimal for *in vivo* transfection, and siRNA/miRNA is effectively delivered and introduced into the cells.

siRNA/Atelocollagen Complex Formation



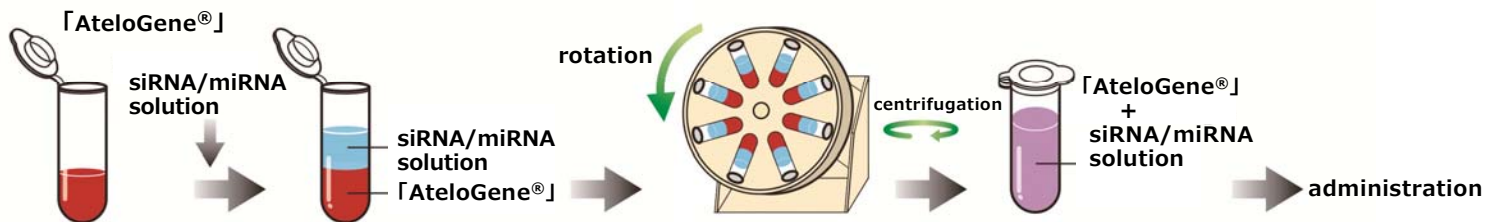
Two types of AteloGene® are available depending on your target tissue or method of administration. **AteloGene® Local Use** is designed for localized administration due to its gelation capability. Gelled siRNA/atelocollagen complexes remain at the injection site and siRNA/miRNA is delivered into the cells effectively.

AteloGene® Systemic Use is suitable for systemic administration via tail vein injection because it does not gelate, and siRNA/miRNA is delivered effectively by the bloodstream throughout the whole body.

Simple handling

AteloGene® procedures are simple and easy.

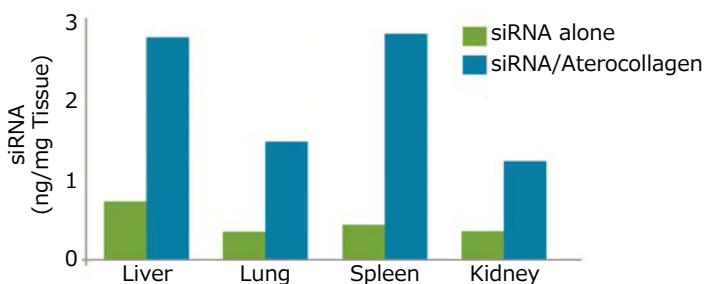
Mix equal volumes of AteloGene® and siRNA/miRNA solution, and administrate the siRNA/miRNA AteloGene® mixture to the mouse.



Efficient Delivery of siRNA/miRNA

AteloGene® enhances delivery of siRNA to tissues.

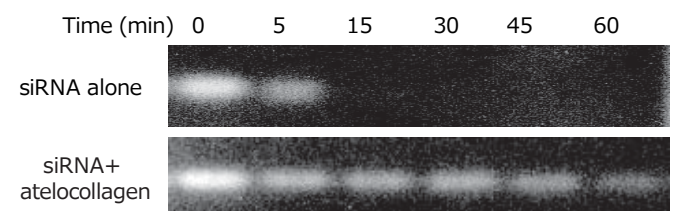
(Takeshita F, *et al.* (2005) *Proc Natl Acad Sci USA*. **102** (34): 12177-12182).



Stability of siRNA/miRNA

siRNA in the atelocollagen/siRNA complex has enhanced stability relative to naked siRNA.

(Minakuchi Y, *et al.* (2004) *Nucleic Acids Res*. **32** (13):e109.)

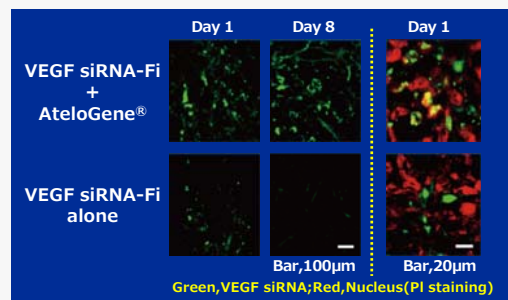


Stability of VEGF siRNA by intratumoral administration

(Data source: Dr. Y. Takei, Nagoya University, Japan)

AteloGene® Local Use was mixed with fluorescent labeled vascular endothelial growth factor (VEGF) siRNA and injected into subcutaneous tumor. Compared to naked siRNA, atelocollagen/siRNA complex gives enhanced effectiveness of transfection, leading to prolonged effectiveness of siRNA treatment.

(Takei Y, et al. (2004) *Cancer Res.* **64** (10): 3365-3370).

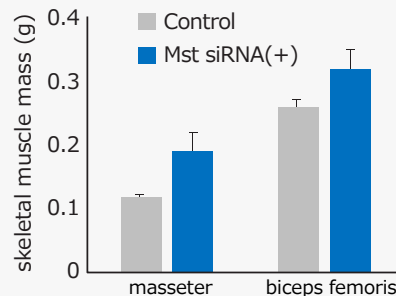


Muscular mass in mice increased by local and systemic administration of myostatin-targeting siRNA

(Data source: Dr. S. Noji, Tokushima University, Japan)

Using the AteloGene® Local Use kit, administration of the atelocollagen/siRNA complex increased molecular mass compared to naked siRNA control. Similar results were seen after systemic administration using AteloGene Systemic Use.

(Kinouchi N, et al. (2008) *Gene Ther.* **15** (15): 1126-1130).

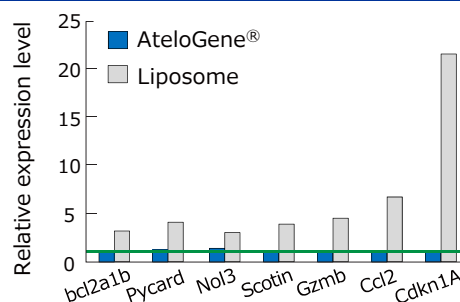


Low Toxicity

Microarray Analysis Shows Low Toxicity of AteloGene®

Expression analysis after administration of siRNA using AteloGene® shows reduced expression of apoptosis-related genes relative to administration of siRNA using a liposomal method.

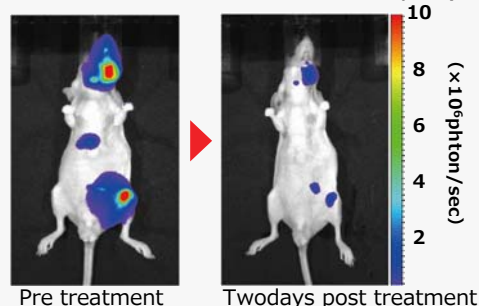
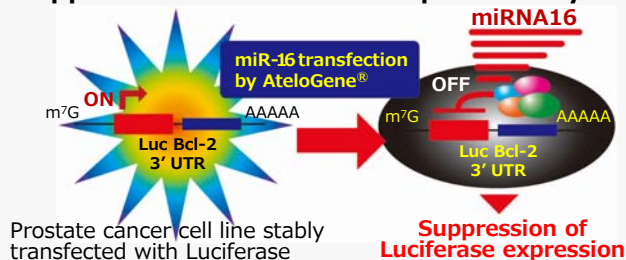
(Ogawa S, et al. (2011) *J Toxicol Sci.* **36** (6): 751-762.)



Systemic delivery of miRNA

(Data source: Dr. F. Takeshita and Dr. T. Ochiya, National Cancer Center Research Institute, Japan)

Suppression of Luciferase expression by miRNA



This reporter system was stably introduced into a metastatic model of prostate cancer cells. Luciferase expression was markedly suppressed when miR-16 was administered systemically with AteloGene® Systemic Use. Marked inhibition of metastasis was also observed. (Takeshita F, et al. (2010) *Mol Ther.* **18** (1): 181-187).

Catalog No.	Product	Quantity	Storage
KKN-1394	AteloGene® Local Use	1 Kit*	2-10°C
KKN-1395	AteloGene® Systemic Use	1 Kit*	2-10°C

*Sufficient for 10 injections. **Agreement is needed to purchase AteloGene®.

For research use only. Not for diagnostic use.



E-mail
info_en@reprocell.com

Headquarters

ReproCELL, Inc.
KDX Shin-yokohama 381 Bldg 9F
3-8-11, Shin-yokohama, Kohoku-ku,
Yokohama, Kanagawa 222-0033, Japan
Tel: +81 (0) 45 475 3887
Fax: +81 (0) 45 474 1006

North America

ReproCELL USA, Inc.
24 Denby Rd. Suite 220
Boston, MA 02134
Tel: +1-617-987-2015
Fax: +1-617-507-2452