A TRIAL OF GENE TRANSFER AUGMENTED BY RADIAL SHOCK WAVE FOR RABBIT CHONDROCYTES IN VITRO

Ryo Murata* Koichi Nakagawa* Seiji Ohtori* Nobuyasu Ochiai* Takahisa Sasho* Yuichi Wada** Hideshige Moriya* * Department of Orthopaedic Surgery, Graduate School of Medicine Chiba University, Chiba City, Japan ** Department of Orthopaedic surgery, Teikyo University Ichihara Hospital, Chiba, Japan

BACKGROUND

• Safety and reproducibility as well as transfection efficiency are required for successful gene therapy.

		Efficiency	Safety	Facility
Viral		0	Δ*	Δ
Non- viral	Ultrasound	Δ~O**	0	0
	Electroporation	Δ	Δ***	Δ
	Gene Gun	Δ	0	Δ

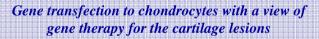
: Toxicity, possibility of malignant mutation

: Augmentation by microbubble agent (Optison® etc)

: Damage of target tissue

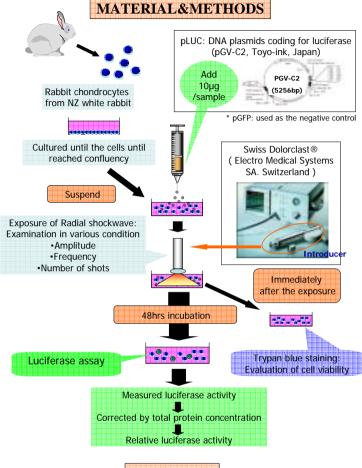
Can extracorporeal shockwave be applied for gene transfer?

- Degenerative or traumatic cartilage lesions are still difficult to treat despite of many conventional treatments
- Gene therapy in the orthopaedic field has not been established

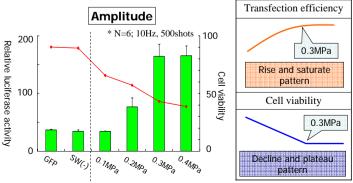


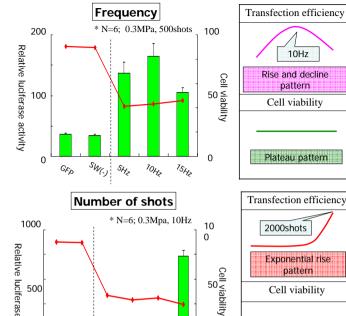
PURPOSE

- Achieve gene transfer with high transfection efficiency for chondrocytes utilizing newly developed radial extracorporeal shockwaves
- Examine the parameters of exposure condition (amplitude, frequency, number of shots) that affect transfection efficiency
- Evaluate cell toxicity of radial extracorporeal shockwaves



RESULTS





DISCUSSION&CONCLUSION

pattern

Cell viability

Plateau pattern

• Luciferase coding DNA plasmid was successfully transfected to rabbit chondrocytes by radial extracorporeal shockwaves.

1000shots

- The transfection efficacy of the exposed cells increased up to about 20-fold compared to the control under the specific condition (0.3 MPa, 2000 shots, 10 Hz).
- The effect of following parameters on the transfection efficiency and the cell viability was investigated.

500shots

200shots

Amplitude: A threshold (nearby 0.3MPa)

500

activity

Frequency: No significant effect on the both factors Number of shots: Dose-dependent effect on the transfection efficiency but not on the cell viability?

- → Each parameter may play the different role on the both factors.
- The transfection efficiency may be augmented by varying these parameters; e.g. more large number of shots.
- Gene transfer by radial shockwaves could be applied for the treatment of various cartilage lesions.