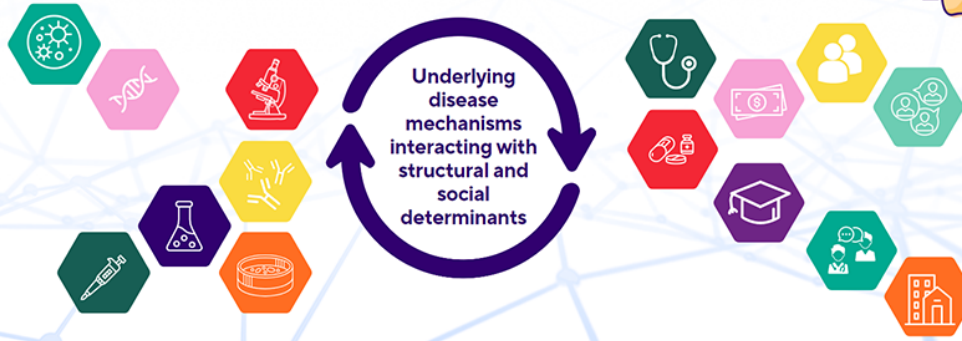




19TH ANNUAL CHILD HEALTH RESEARCH DAYS
Outcomes in Child Health



October 25 + 26, 2023 | RBC Convention Centre, Winnipeg, Manitoba

Abstract Submission Form

CHRD 2023: Abstract Submission Form

Submitter Name

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Presenter Name

Yuichiro Miyake

Presenter Status

Post-Doctoral Fellows

Research Category

Basic Science

Role in the project

Design
Perform Experiments
Analyze Data
Write Abstract

Title

Using microinjection for drug delivery with nanoparticles in lung explants to study prenatal treatment of congenital diaphragmatic hernia.

Background

Congenital diaphragmatic hernia (CDH) is associated with abnormal lung development resulting in lung hypoplasia and pulmonary hypertension, and fetoscopic endoluminal tracheal occlusion (FETO) improves the survival rate in fetuses with severe CDH. To further improve outcomes, microinjection into the trachea for drug delivery has potential as a clinical therapy. Here, we present an ex vivo microinjection technique with rat lung explants to study prenatal therapy with nanoparticles (NPs).

Objective

We used microsurgery to isolate lungs from rats on embryonic day 18. We injected chitosan nanoparticles loaded with fluorescein (FITC) into the trachea of the lung explants.

Methods

We compared the difference in biodistribution of two types of nanoparticles, functionalized IgG-conjugated nanoparticles (IgG-nanoparticles) and bare nanoparticles after 24 hours culture with immunofluorescence (IF). We used IF to mark lung epithelial cells with E-cadherin and to investigate an apoptosis (Active-caspase 3) and inflammatory marker (Interleukin, IL-6) and compared its abundance between the two experimental groups and control lung explants.

Results

Lung explant samples came from two different litters for each group. We detected the presence of nanoparticles in the lung explants, and the relative number of nanoparticles to cells was 2.49 fold higher in IgG-nanoparticles than bare nanoparticles ($p < 0.001$). Active caspase-3 protein abundance was similar in the control, bare nanoparticles (1.08 fold), and IgG-nanoparticles (1.47 fold) groups ($p = 0.34$). Similarly, IL-6 protein abundance was not different in the control, bare nanoparticles (1.13 fold), and IgG-nanoparticles (1.07 fold) groups ($p = 0.48$).

Conclusion

Our novel microinjection model to deliver nanoparticles effectively into the trachea with rat lung explants holds promising potential as a tool to study prenatal treatment for CDH, and provides insights into a new prenatal therapy. This model offers a controlled platform to investigate the potential of prenatal therapy with nanoparticles.

Table/Figure File

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Nano particle (FITC)

E-Cadherin

DAPI

Merged

Bare CNP

IgG- CNP

