

# human induced pluripotent stem cells derived airway epithelium

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#### Organisation

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#### SCOPE OF THE METHOD

The Method relates to	Human health
The Method is situated in	Translational - Applied Research
Type of method	In vitro - Ex vivo

#### DESCRIPTION

#### Method keywords

Human induced Pluripotent Stem Cell

differentiation airways lung epithelial cells

#### Scientific area keywords

Induced pluripotent stem cells human airways 3D organoid models Chronic obstructive pulmonary disease asthma epithelial cells co-culture

#### Method description

We devised a simple and reliable method for reprogramming peripheral blood mononuclear cells into hiPSC and then to differentiate them into air-liquid interface bronchial epithelium (iALI) within 45 days. Of note, this method does not involve any cell sorting step. We reprogrammed blood cells from one healthy control and three patients with very severe COPD. The mean cell purity at definitive endoderm and ventral anterior foregut endoderm was >80%, assessed by CXCR4 and NKX2.1 expression respectively. vAFE cells from all four hiPSC differen-tiated into bronchial epithelium in air-liquid interface (ALI) conditions, with large zones covered by beating ciliated, basal, goblets, club cells and neuroendocrine cells, as found *in vivo*.

#### Lab equipment

Flow cytometry, Biosafety cabinet, Microscopy phase contrast, Fluorescence PCR, Matrix (geltrex, matrigel), Transwell insert for culture, Transepithelial/transendothelial electrical resistance.

#### Method status

History of use

Internally validated Published in peer reviewed journal

## PROS, CONS & FUTURE POTENTIAL

#### Advantages

Robust, efficient and reproducible protocol Human normal lung development modeling and diseases modeling Stem cells: renewable and sustainable source of airway epithelial cells High-Yield Human Induced Pluripotent Stem Cell-Derived airway epihelial cells Personalized medicine High input pharmacological screening Genome editing CRISPR Cas9 technology

### Challenges

Cost,

Mandatory to check regulary genetic stability of stem cells during culture maintenance,

Derivation of clinical grade iPSC culture and derived therapeutic cells.

## Modifications

Optimization of differentiation protocol, Co-culture with other cell type such as immune cells

#### Future & Other applications

Disease modeling cell therapy

## **REFERENCES, ASSOCIATED DOCUMENTS AND OTHER INFORMATION**

## References

Differentiation protocol frome the team PMID: 35954266 iPSC cell lines derived PMID: 34624616 , PMID: 33099111, PMID: 30296669 Kotton DN protocol lab: PMID: 35781291, PMID: 35499347 Gotoh Lab: PMID: 34798066

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