

Tissue-Engineered Lung: An In-Vivo Model of Airway Injury and Regeneration

Denise Al Alam, PhD

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Background

- Lung disease causes more than 230,000 deaths per year in the US
- Human lung often responds to injury by scarring rather than repair
- Molecular and cellular mechanisms of human lung regeneration are largely unknown
- Currently existing models of lung regeneration are somewhat limited

GOAL

To develop a more “complete” model of in vivo lung regeneration that allows to study the complex mechanisms driving lung regeneration

Methods

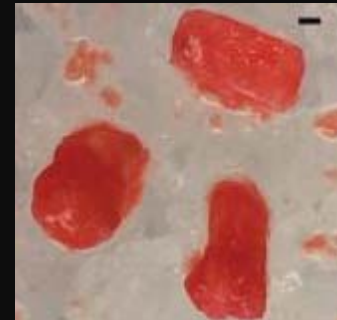


Native postnatal lung
(14 days)

Dissociation



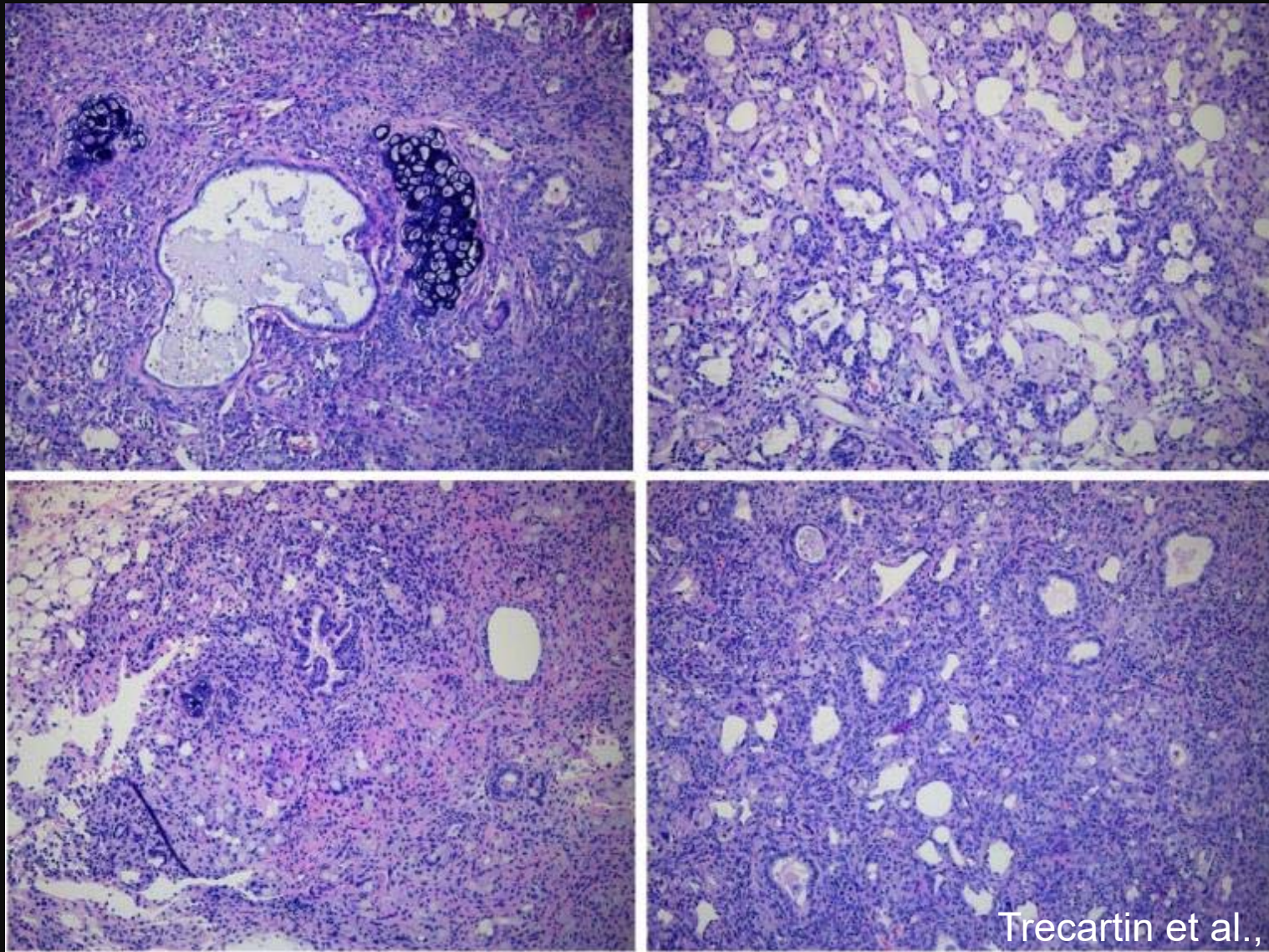
Collect at different time points



Seeded polymers

Trecartin et al., 2016

Distinguishable proximal and distal structures via H&E

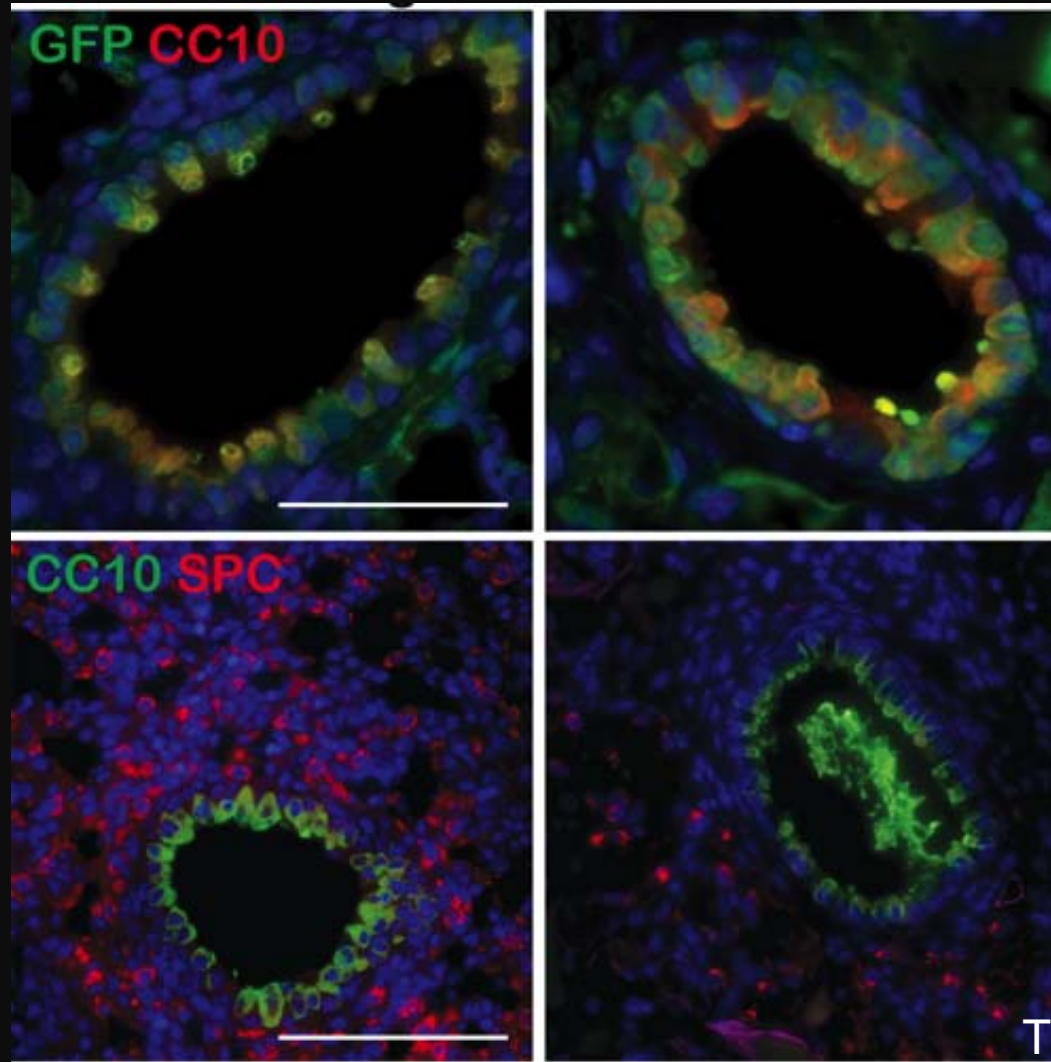


TELu airways are of donor origin

GFP native lung implanted into NOD-SCID mouse for 4 weeks

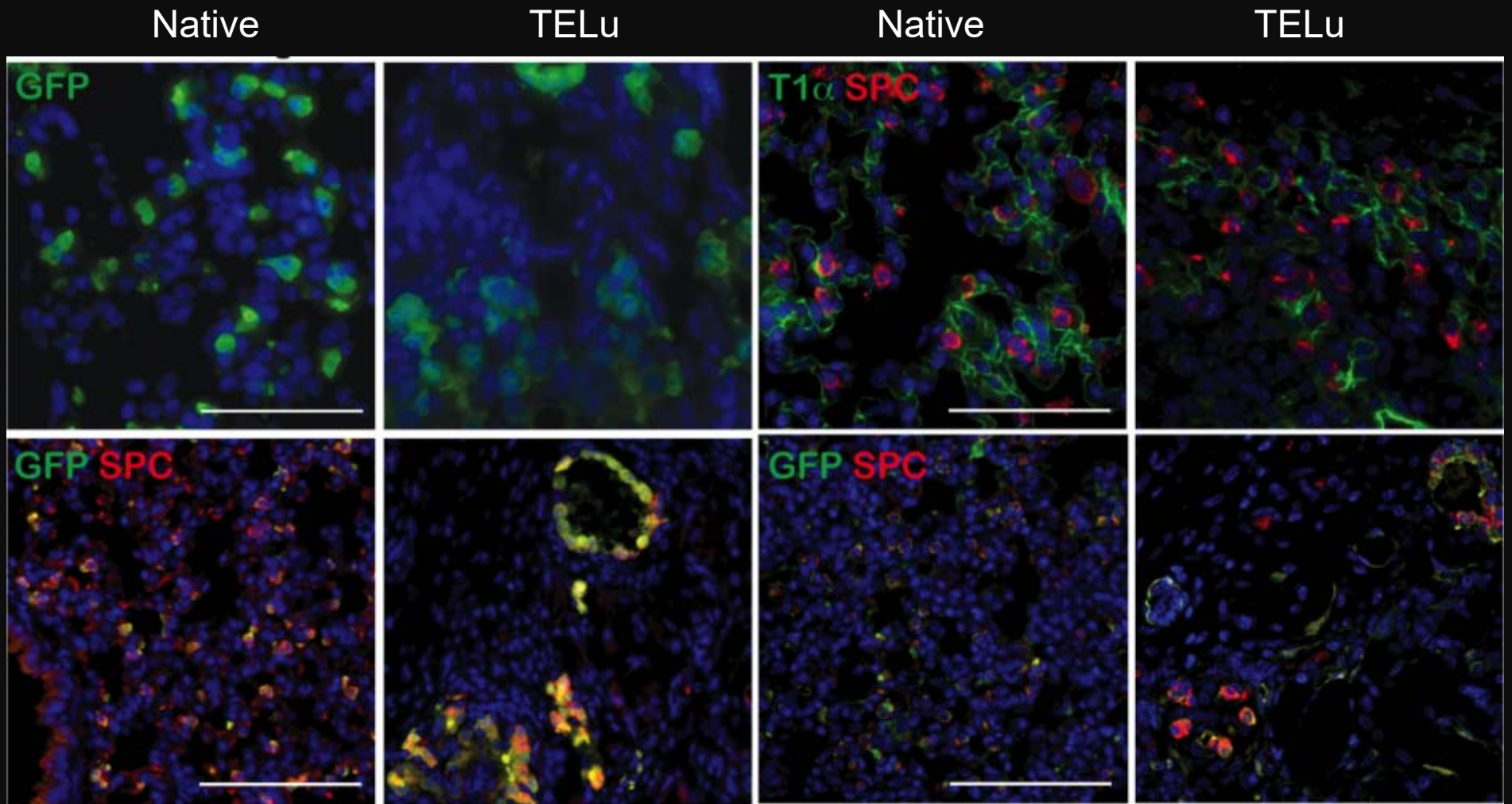
Native

TELu

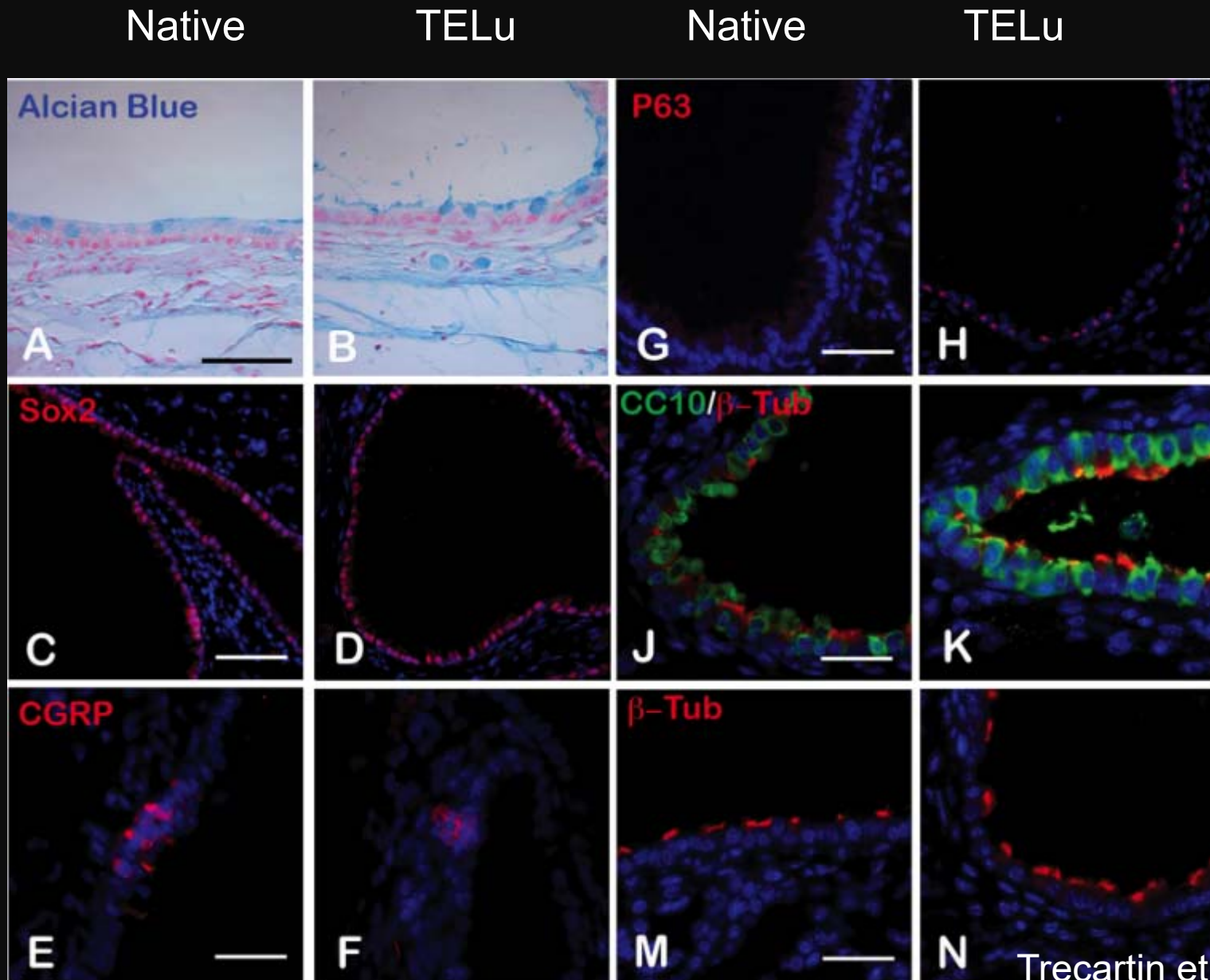


Trecartin et al., 2016

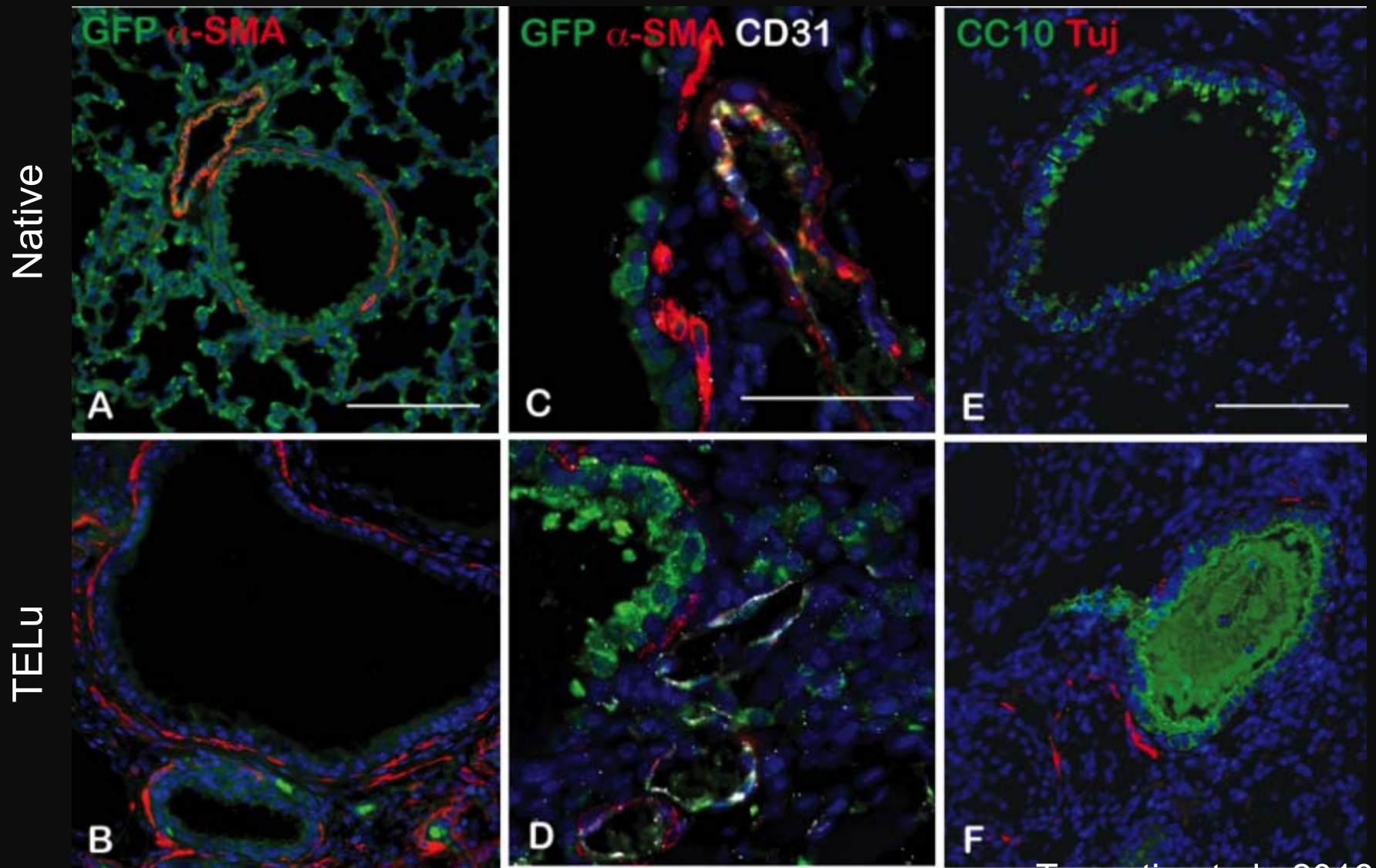
SPC positive cells are derived from donor cells



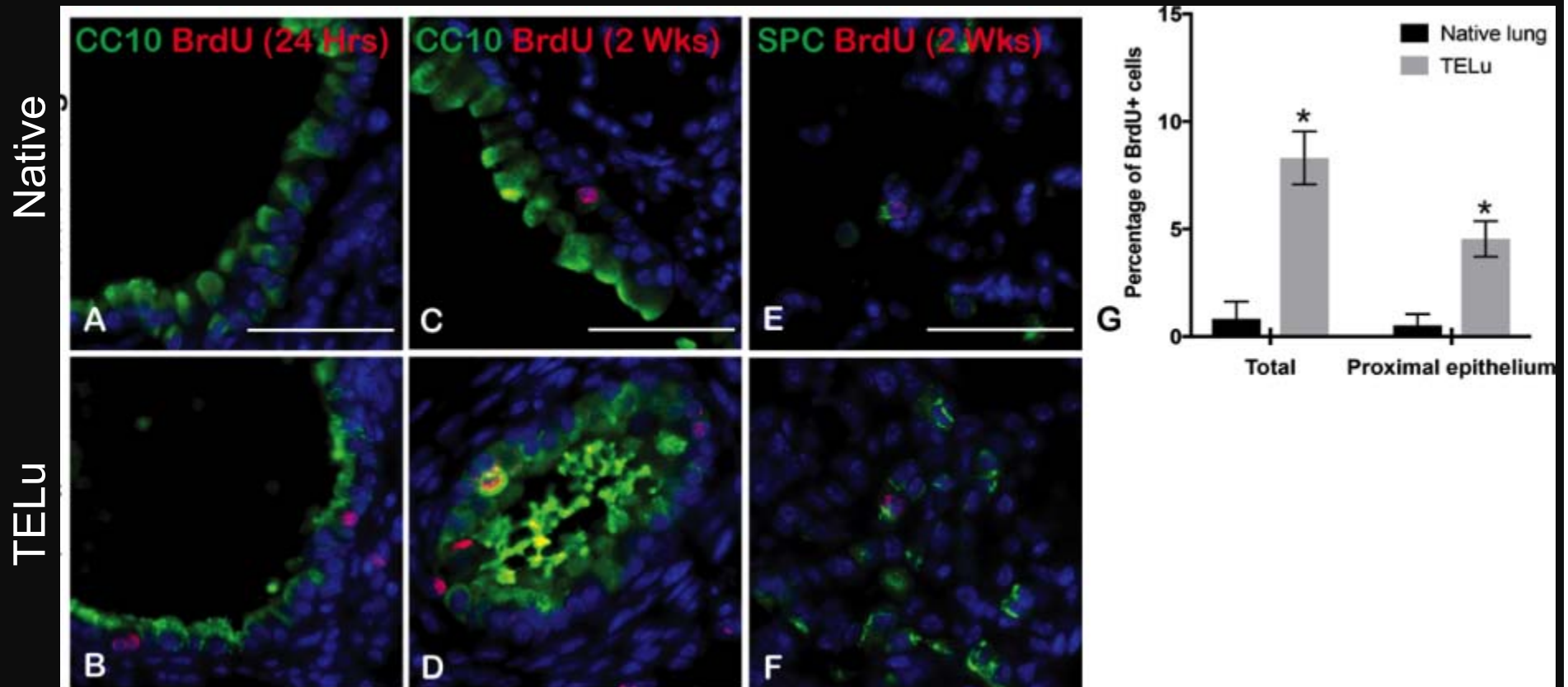
Characterization of epithelial markers



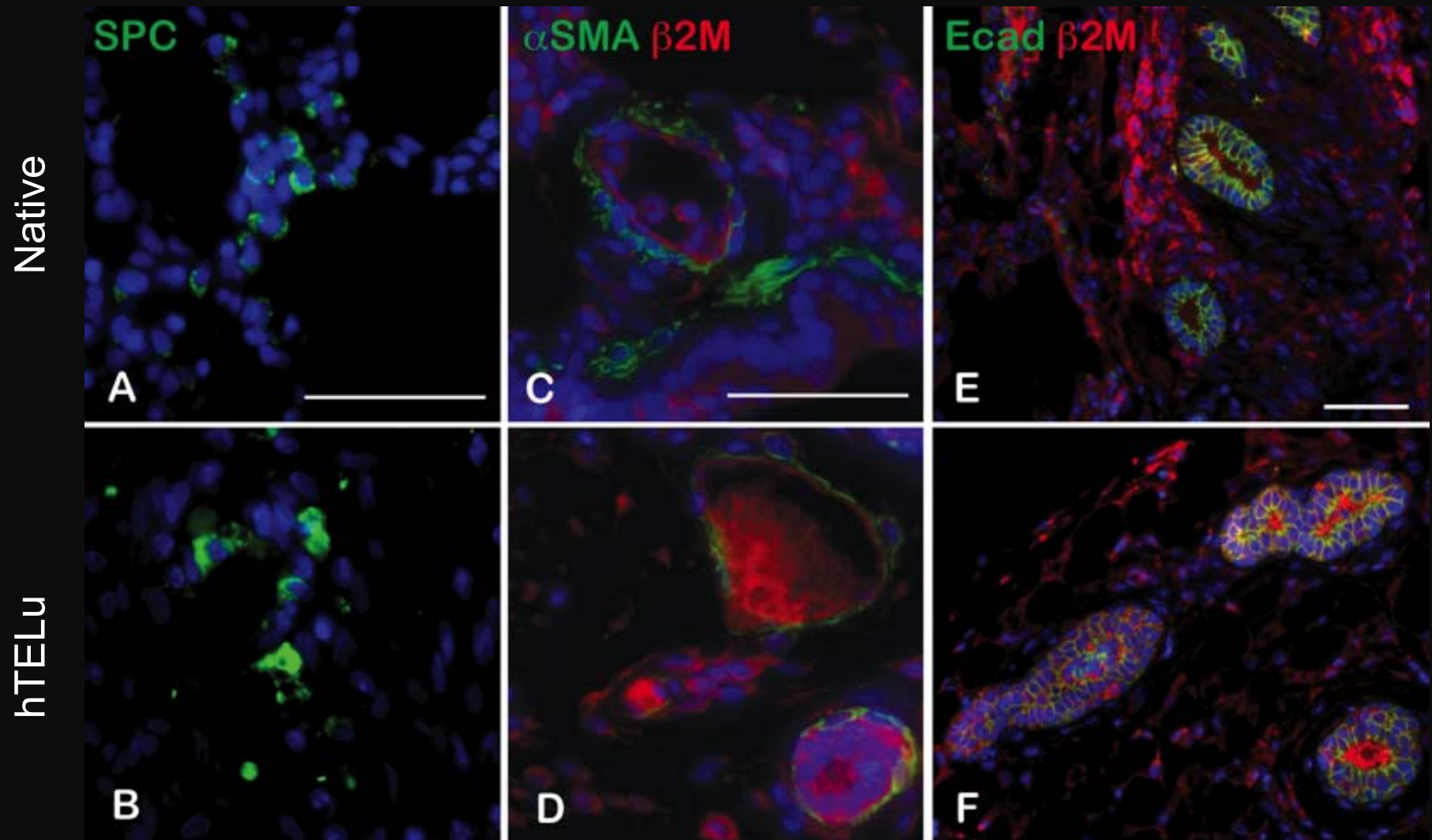
Characterization of mesenchymal markers



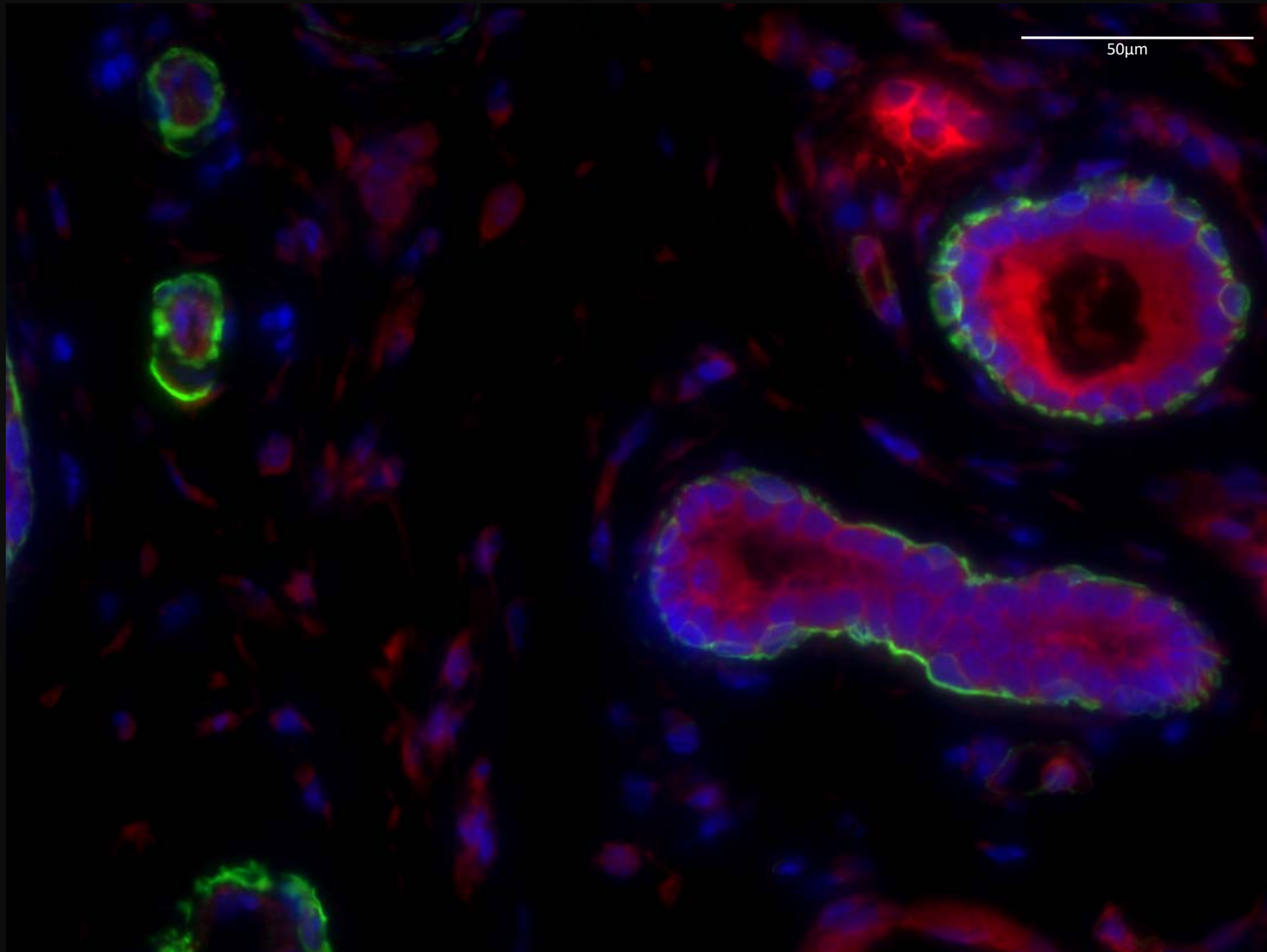
Higher proliferative rate seen in TELu



Generating Human TELu

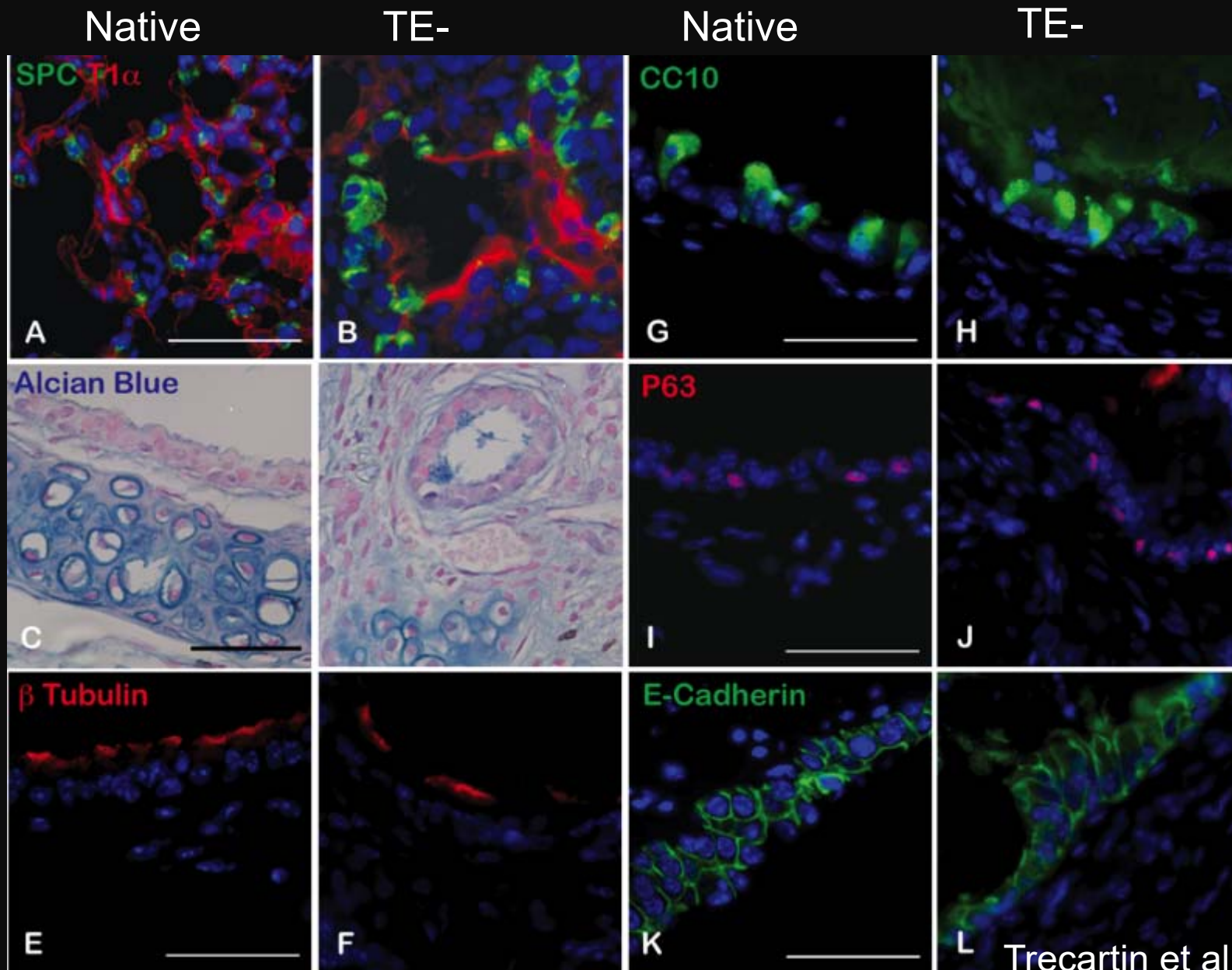


Smooth Muscle cells in hTELu



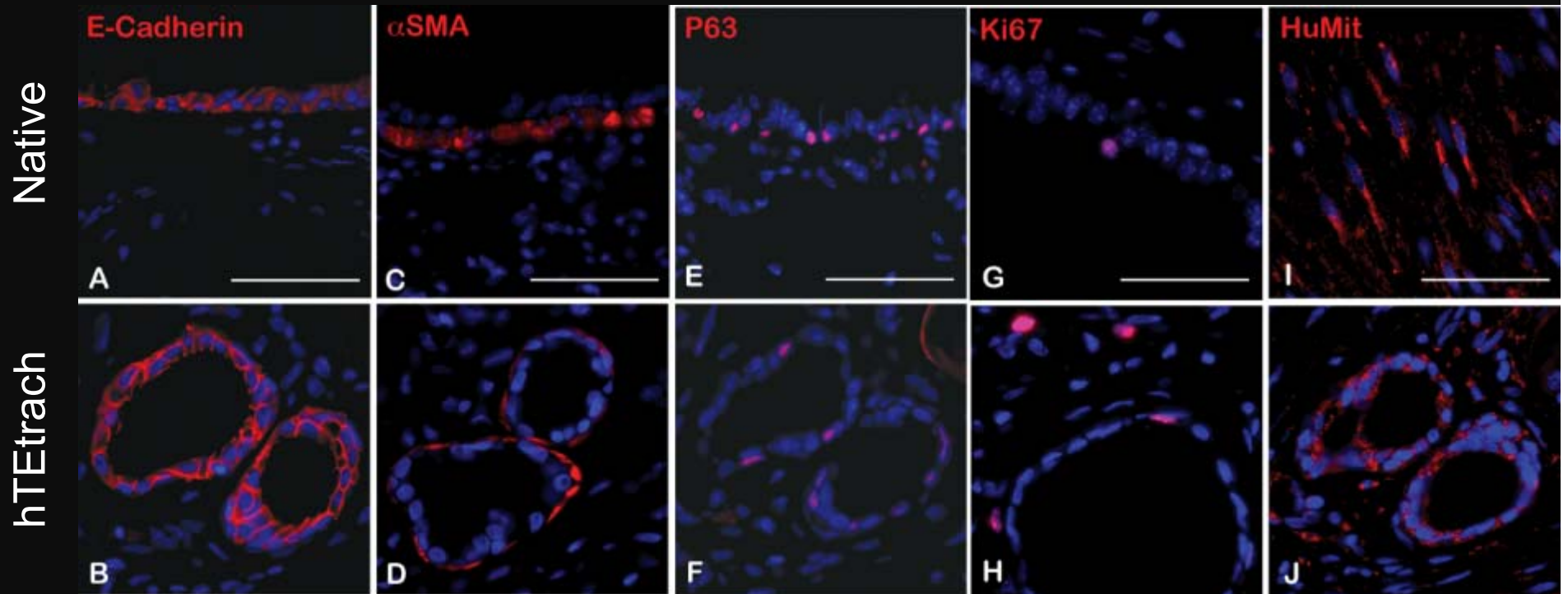
α -SMA B2-micro

Generation of compartment specific tissue

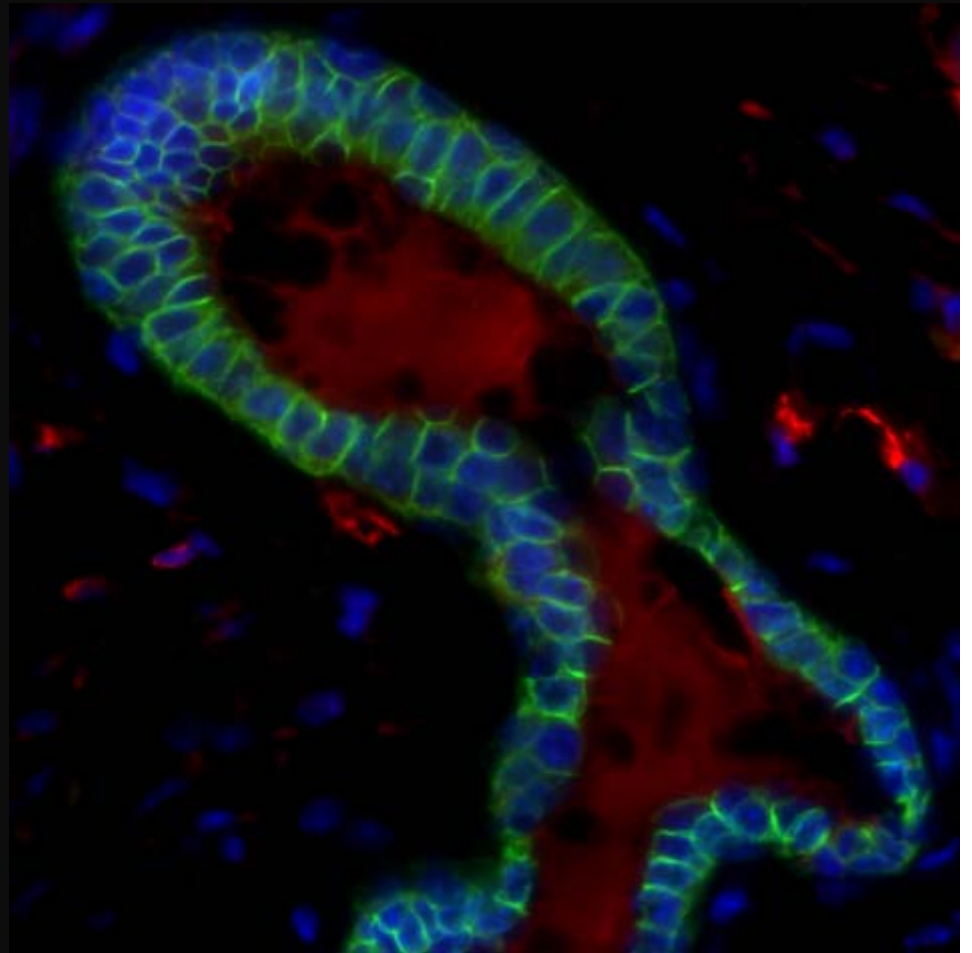
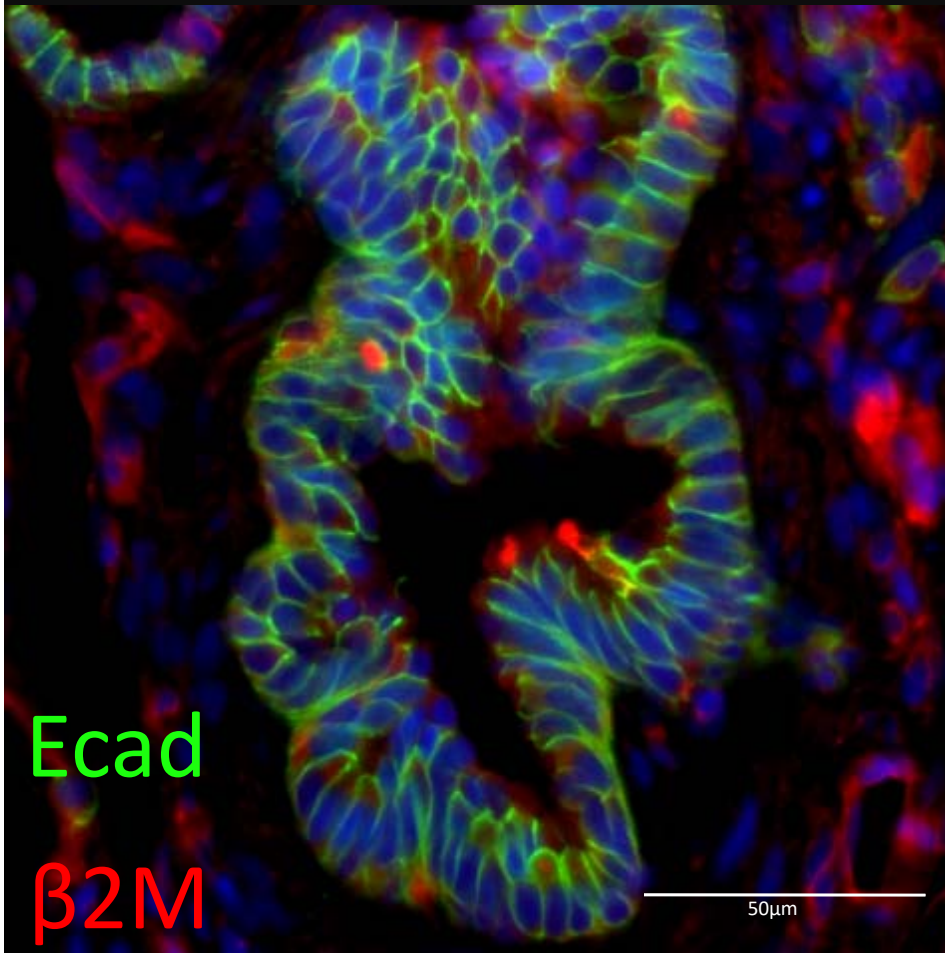


Trecartin et al., 2016

Human TE-Trach displaying epithelial and mesenchymal markers



hTELu Contains Large Epithelial Structures



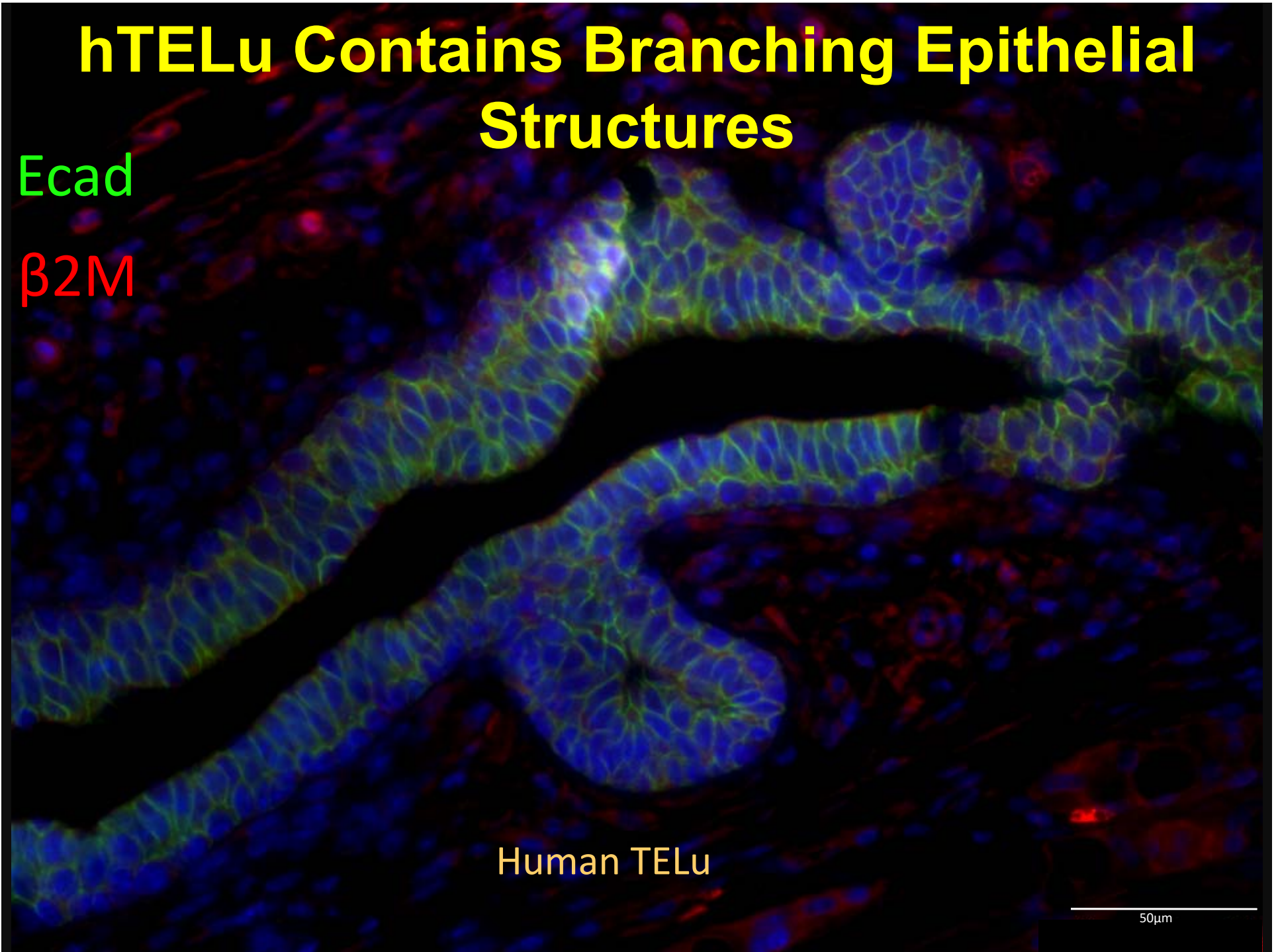
hTELu Contains Branching Epithelial Structures

Ecad

β 2M

Human TELu

50 μ m



Summary

- Murine TELu containing all cell types, e.g. SPC positive cells, likely AEC2s, Club cells, Ciliated cells, Smooth muscle cell and nerve cells
- Human TELu containing epithelial structures, SPC positive cells, Smooth muscle cells
- Proximal and distal regional structures can be generated distinctly

ACKNOWLEDGMENTS

A photograph of the Children's Hospital Los Angeles building at dusk. The building is a large, modern structure with a glass facade that reflects the warm colors of the sunset. The hospital's name and logo are visible on the upper part of the building. In the foreground, there are palm trees and a parking lot with several cars. The sky is a mix of purple, pink, and blue.

Grikscheit Lab:

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