# MICHIGAN STATE UNIVERSITY

## INTRODUCTION

- Pulmonary Hypertension (PH) is a complex disorder associated with elevated mean pulmonary arterial pressure (>25 mmHg at rest) and progressive pathologies in pulmonary arteries.
- Increased resistance and stiffness in vessels, stiffened Pulmonary Artery (PA).
- Right ventricle increased after-load
- Left untreated, the median survival: 2.8 years

#### **Objectives**



• Develop a model to account for the **micro-structural stiffness** in PA and represents the overall stiffness in macro scale for each constituent

# METHODS

### **Clinical Data**

- Clinical data was prospectively acquired in four PH subjects (UM1-2) and UM5-6) and a control subject (CT1) with cardiac transplant with pulmonary normotensive response.
- 3D data on vascular anatomy were acquired using cine MRI. PC-MRI to obtain dynamic data for luminal area changes and blood flow velocity.

## **Finite Element Analysis**

- 3D anatomy is meshed and the finite element is performed.
- Constrained-mixture theory for three constituent of elastin, collagen and **smooth muscle cells** is adopted.

$$w_{e} = \frac{c}{2} (\mathbf{C} : \mathbf{I} - 3)$$

$$w_{c} = \sum_{i=1}^{4} \phi_{i}^{c} \frac{k_{1}^{i}}{4k_{2}^{i}} \left[ \exp\left[\frac{k_{2}^{i}(\lambda_{k}^{2} - 1)^{2}\right] - 1 \right]$$

$$w_{m} = \frac{m_{1}}{4m_{2}} \left[ \exp\left[\frac{m_{2}(\lambda_{k}^{2} - 1)^{2}\right] - 1 \right] + w_{a}$$

$$\mathbf{F}_{\alpha} = \mathbf{G}_{\alpha} \mathbf{F}$$

## Optimization

• The adjoint-based method is employed for optimizing the vascular constituent parameters.

$$\min_{\mathscr{P}} J \Big|_{P} = \frac{1}{2} \big| D - D_{c} \big|^{2} \qquad \frac{dJ}{d\mathscr{P}} = - \left[ \frac{\partial F}{\partial \mathscr{P}} \right]^{T}$$



# **Patient-specific in-vivo Arterial Characterization in Pediatric Pulmonary Hypertension**

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## RESULTS

#### **Optimized Parameters**

- for three sets of low, middle and high pre-stretch G.
- stresses for the patients are significantly higher than the normotensive subject.

## **Stress and Stiffness Analysis**

- two constituent.
- PH subjects.
- as the other two stiffens.

# DISCUSSION

- stiffness.
- behavior is activated when the pressure is high.
- by age.

# REFERENCES

1.Wang, Z., & Chesler, N. C. (2011). Pulmonary vascular wall stiffness: an important contributor to the increased right ventricular afterload with pulmonary hypertension. Pulmonary circulation, 1(2), 212-223.

2. Baek, S., Rajagopal, K. R., & Humphrey, J. D. (2006). A theoretical model of enlarging intracranial fusiform aneurysms.

3. S.M. Shavik, Z. Jiang, S. Baek, L. C. Lee, High-resolution multi-organ finite element modeling of ventricular-arterial coupling, Frontiers in Physiology, vol. 9, 119, 2018

**Project Title: (U01 HL135842)** Image-based Multi-scale Modeling Framework of the Cardiopulmonary System: Longitudinal Calibration and Assessment of Therapies in Pediatric Pulmonary Hypertension

• The optimized parameters are collected for the smallest functionals

• The optimized stiffness parameters ( $c, k_1, m_1$ ) and circumferential

• For all the subjects, elastin carry more load compared to the other

• Stiffness as the change in the nominal stress with respect to the fiber stretch is calculated. For all patients the stiffness has been higher for

• Over a cardiac cycle toward the systole the stiffness of elastin drops

• The elevated stiffness could be attributed to fiber growth, removal and remodeling of different constituents with varying intrinsic

• The dominance of the load-carrying capacity of elastin over the other two constituents could be due to the fact that the PA operates in low range of pressure and collagen and smooth muscle stiffening

• The younger age of the patients even though hypertensive could be a factor to be looked at. The constituent production and removal vary