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Wave Intensity Analysis Provides Novel Insights into Pulmonary Hypertension

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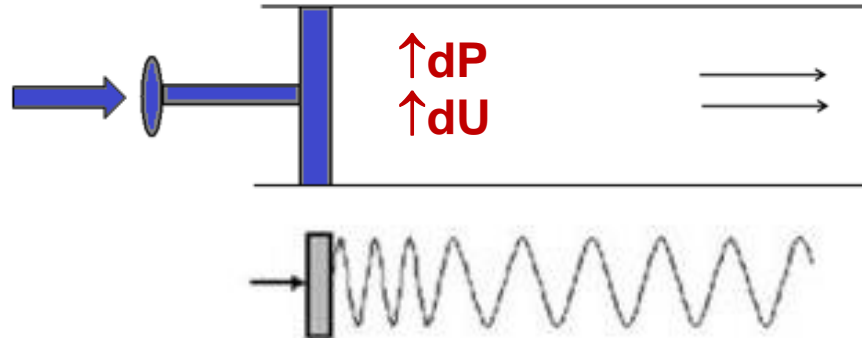


Background

Arterial waves

Proximal artery

Forward compression wave
(FCW)

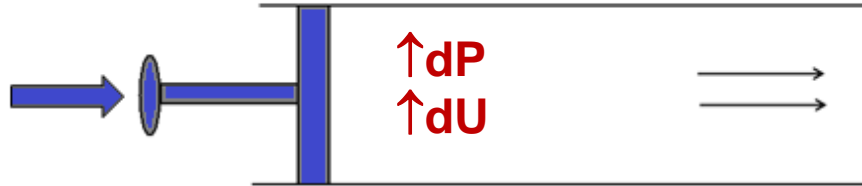


P: pressure
U: velocity

Arterial waves

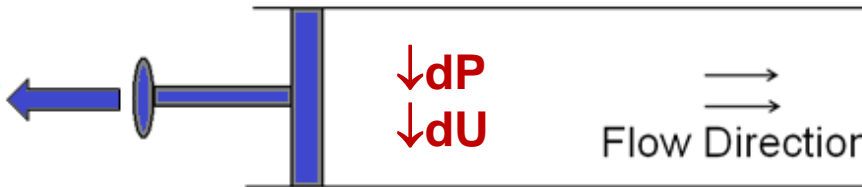
Proximal artery

Forward compression wave
(FCW)



P: pressure
U: velocity

Forward decompression wave
(FDW)



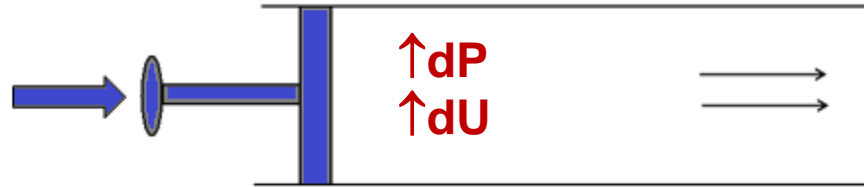
Flow Direction



Arterial waves

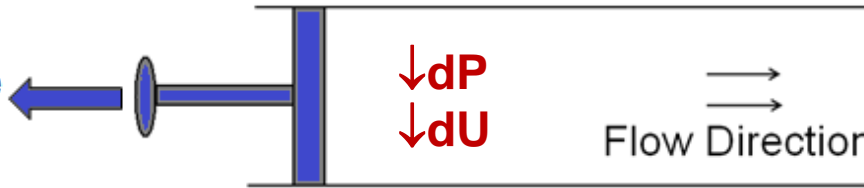
Proximal artery

Forward compression wave
(FCW)



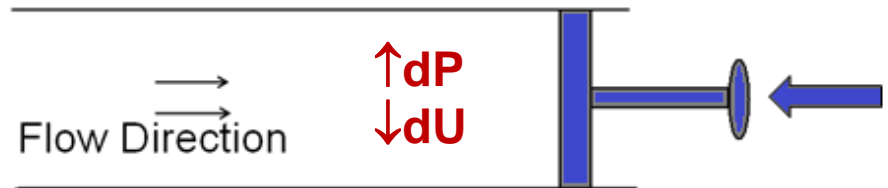
P: pressure
U: velocity

Forward decompression wave
(FDW)



Distal artery

Backward compression wave
(BCW)



Backward decompression wave
(BDW)

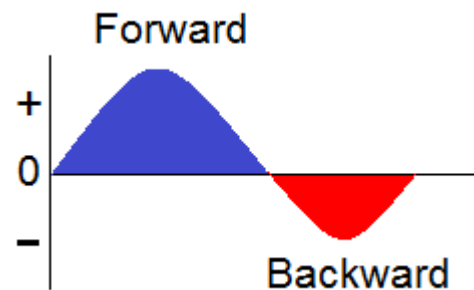


Proximal

Distal

Wave intensity analysis (WIA)

- Analysis of the incremental changes in pressure (P) and flow velocity (U) in a circulation
- Wave intensity: energy carried by a wave
 $dI = dP \times dU$ (Unit $W/m^2 = J/sm^2$)



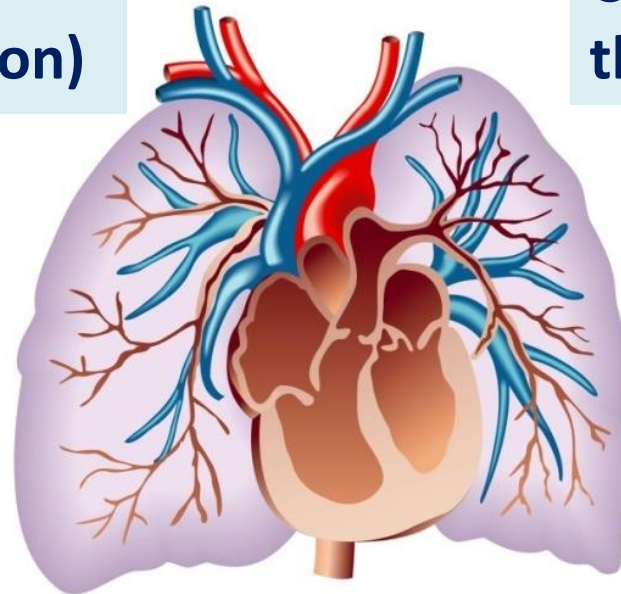
Pulmonary hypertension (PH)

Mean pulmonary arterial pressure (PAPm) ≥ 25 mmHg
→ right heart failure

PAH (Pulmonary
arterial hypertension)

CTEPH (Chronic
thromboembolic PH)

PH due to left
heart disease



PH with unclear or
multifactorial causes

PH due to lung disease

WIA in pulmonary artery



EUROPEAN RESPIRATORY *journal*

OFFICIAL SCIENTIFIC JOURNAL OF THE ERS

Assessment of ventriculo-arterial interaction in pulmonary arterial hypertension using wave intensity analysis

Edmund M.T. Lau^{1,2,3}, David Abelson², Nathan Dwyer⁴, Young Yu^{1,2}, Martin K. Ng^{1,2} and David S. Celermajer^{1,2}



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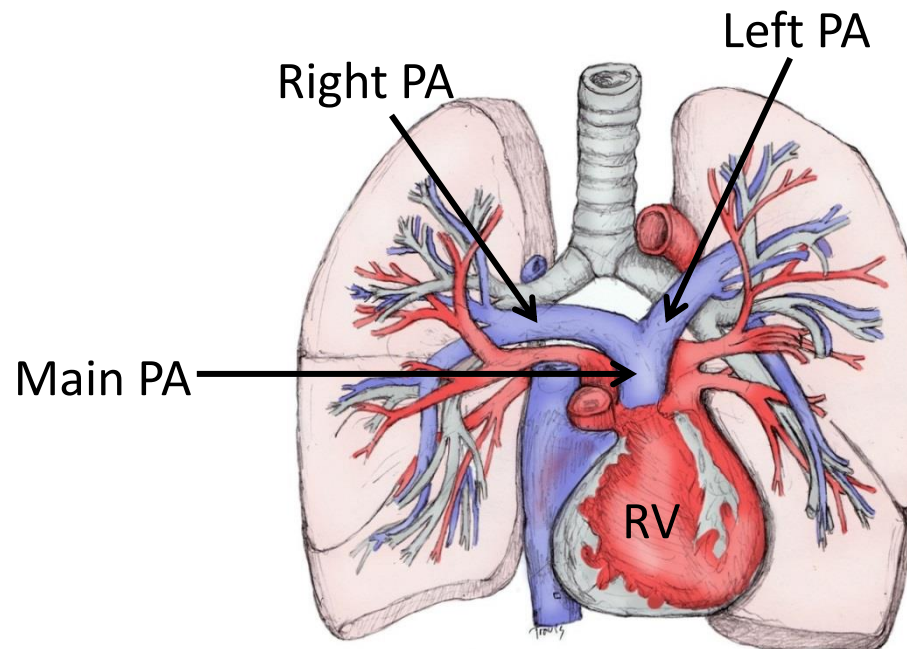
Noninvasive pulmonary artery wave intensity analysis in pulmonary hypertension

Michael A. Quail,¹ Daniel S. Knight,^{1,2} Jennifer A. Steeden,¹ Liesbeth Taelman,³ Shahin Moledina,¹ Andrew M. Taylor,¹ Patrick Segers,³ J. Gerry Coghlan,² and Vivek Muthurangu¹

- **WIA in the pulmonary artery in man is feasible!**
- **Clinical implications?**

Objective

- **Assess arterial wave characteristics in the pulmonary artery**
- **Explore the clinical usefulness of WIA in pulmonary hypertension**





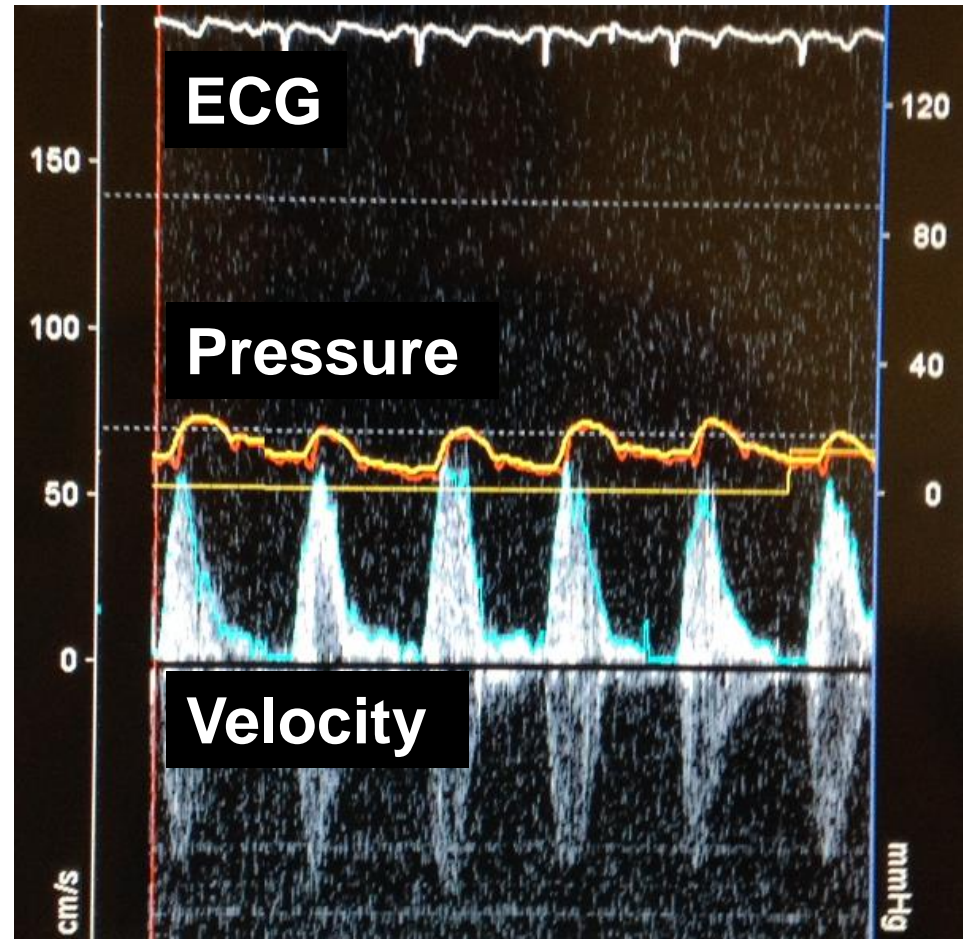
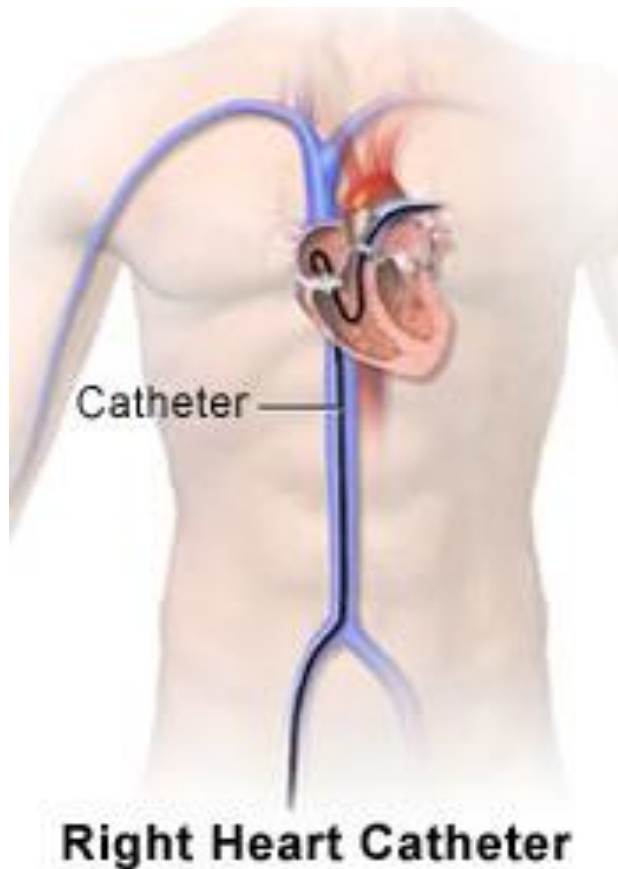
Study design

Inclusion criteria

- **Patients referred to the cardiac catheterisation laboratory for clinical reasons**
- **Control subjects: no significant heart or lung disease**
- **PH patients: patients with confirmed or suspected PAH or CTEPH**

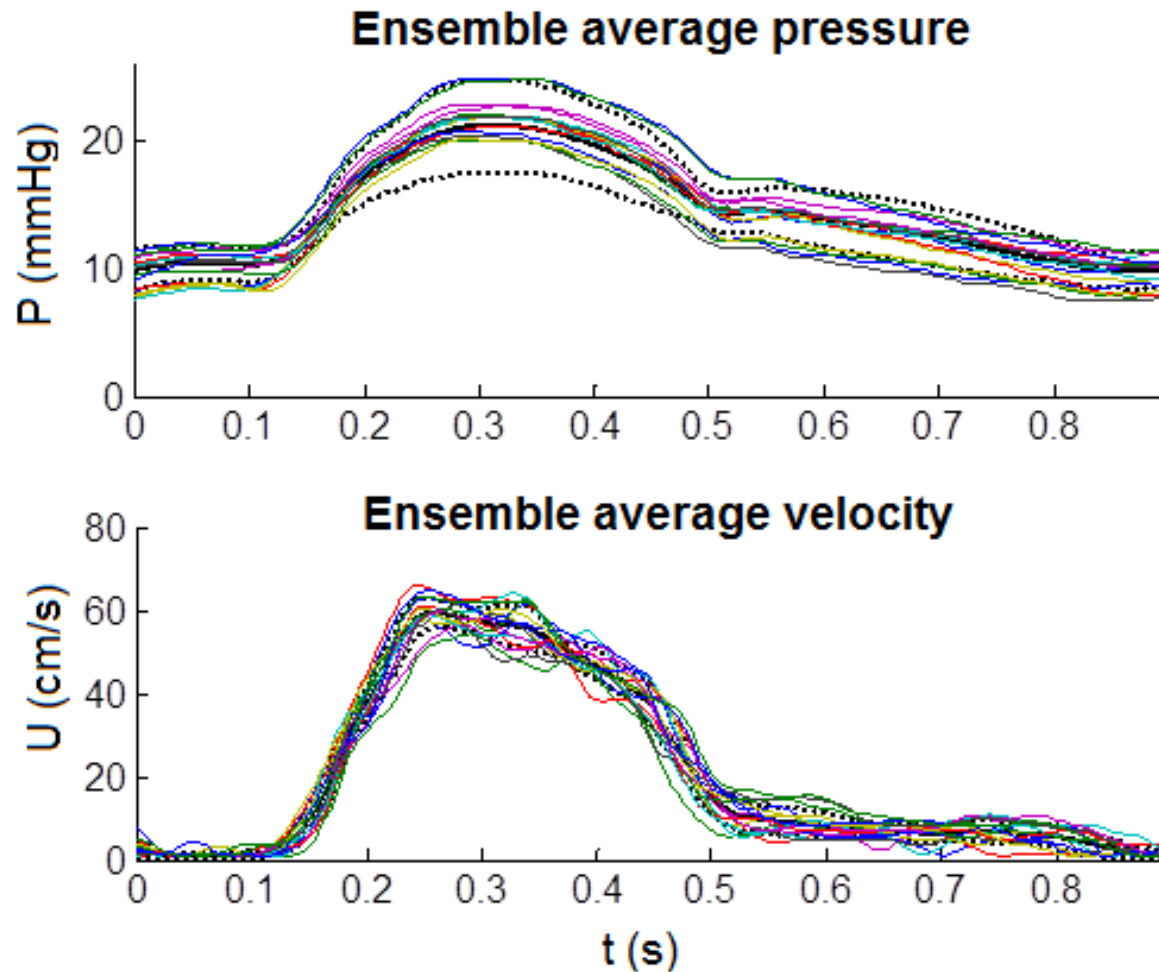
Right heart catheterisation

Right heart catheterisation with simultaneous pressure and velocity measurements.



Data processing

- Data ensemble averaged using the R-wave of ECG



Data processing

- Data ensemble averaged using the R-wave of ECG
- **Calculation of wave speed (sum of squares method)**

$$c = \frac{1}{\rho} \cdot \sqrt{\frac{\sum dP^2}{\sum dU^2}}$$

Data processing

- Data ensemble averaged using the R-wave of ECG
- Calculation of wave speed (sum of squares method)
- **Wave intensity (WI) normalized to number of samples in the cardiac cycle**

$$WI = dP \frac{CCD}{dt} \cdot dU \frac{CCD}{dt}$$

CCD: cardiac cycle duration

Data processing

- Data ensemble averaged using the R-wave of ECG
- Calculation of wave speed (sum of squares method)
- Wave intensity (WI) normalized to number of samples in the cardiac cycle
- **Separation of forward and backward waves**

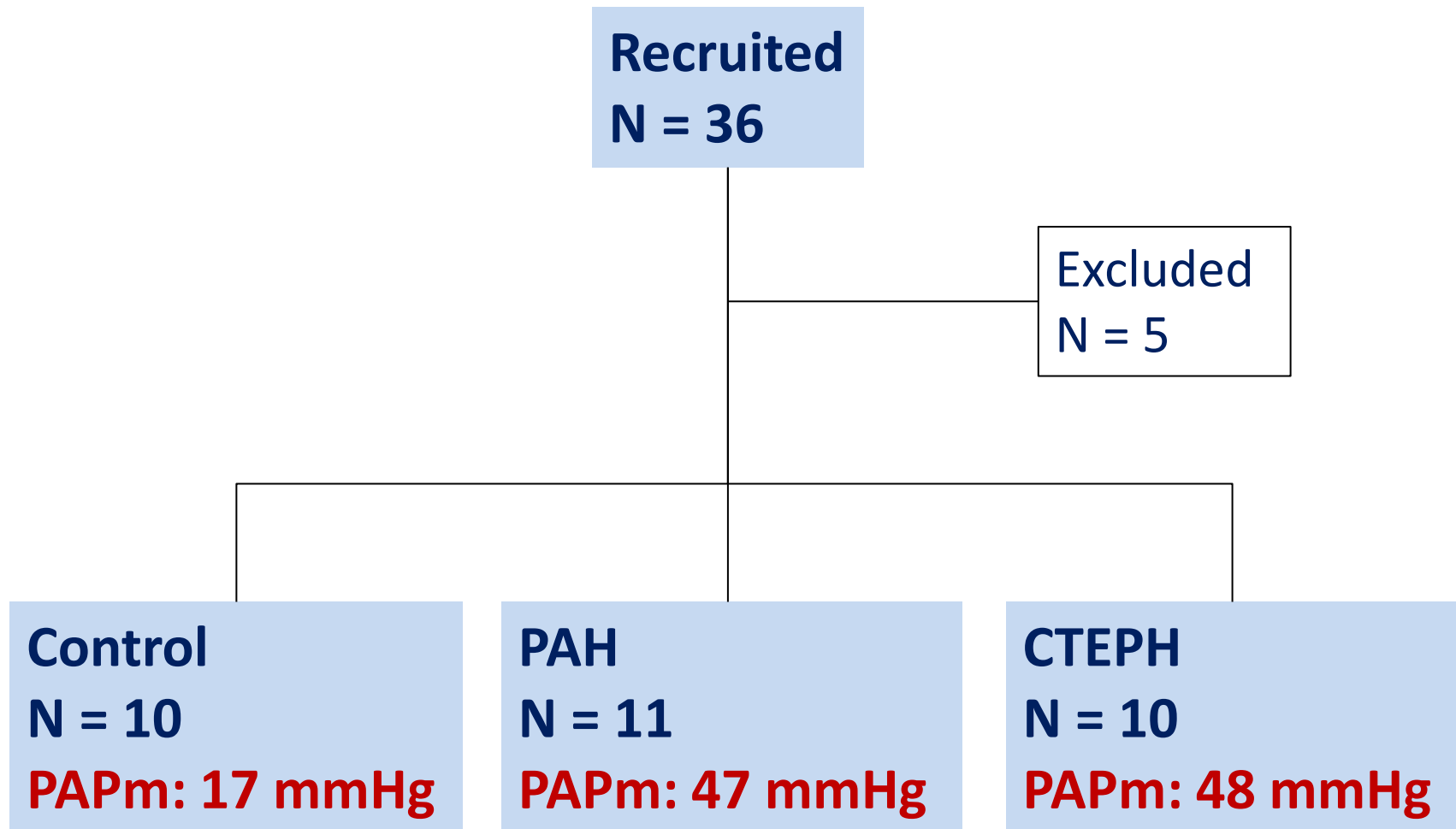
$$\textit{Forward: } WI_+ = \left(\frac{dP \cdot CCD}{dt} + \rho \cdot c \frac{dU \cdot CCD}{dt} \right) / 4\rho c$$

$$\textit{Backward: } WI_- = - \left(\frac{dP \cdot CCD}{dt} - \rho \cdot c \frac{dU \cdot CCD}{dt} \right) / 4\rho c$$



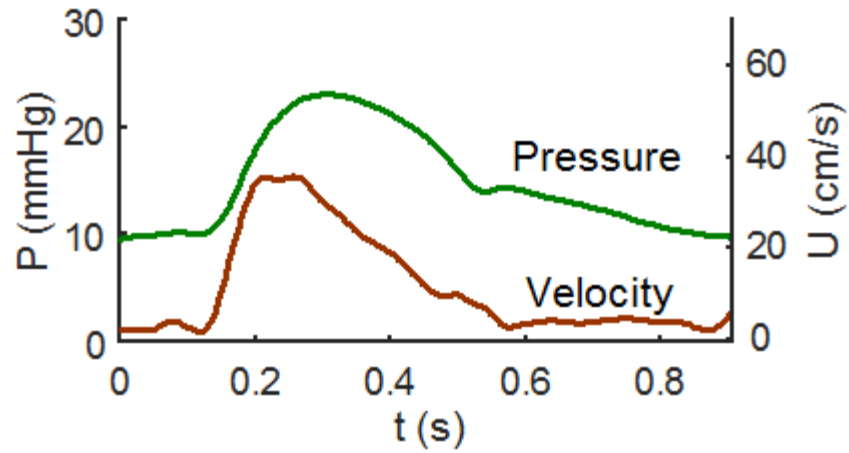
Results

Study subjects

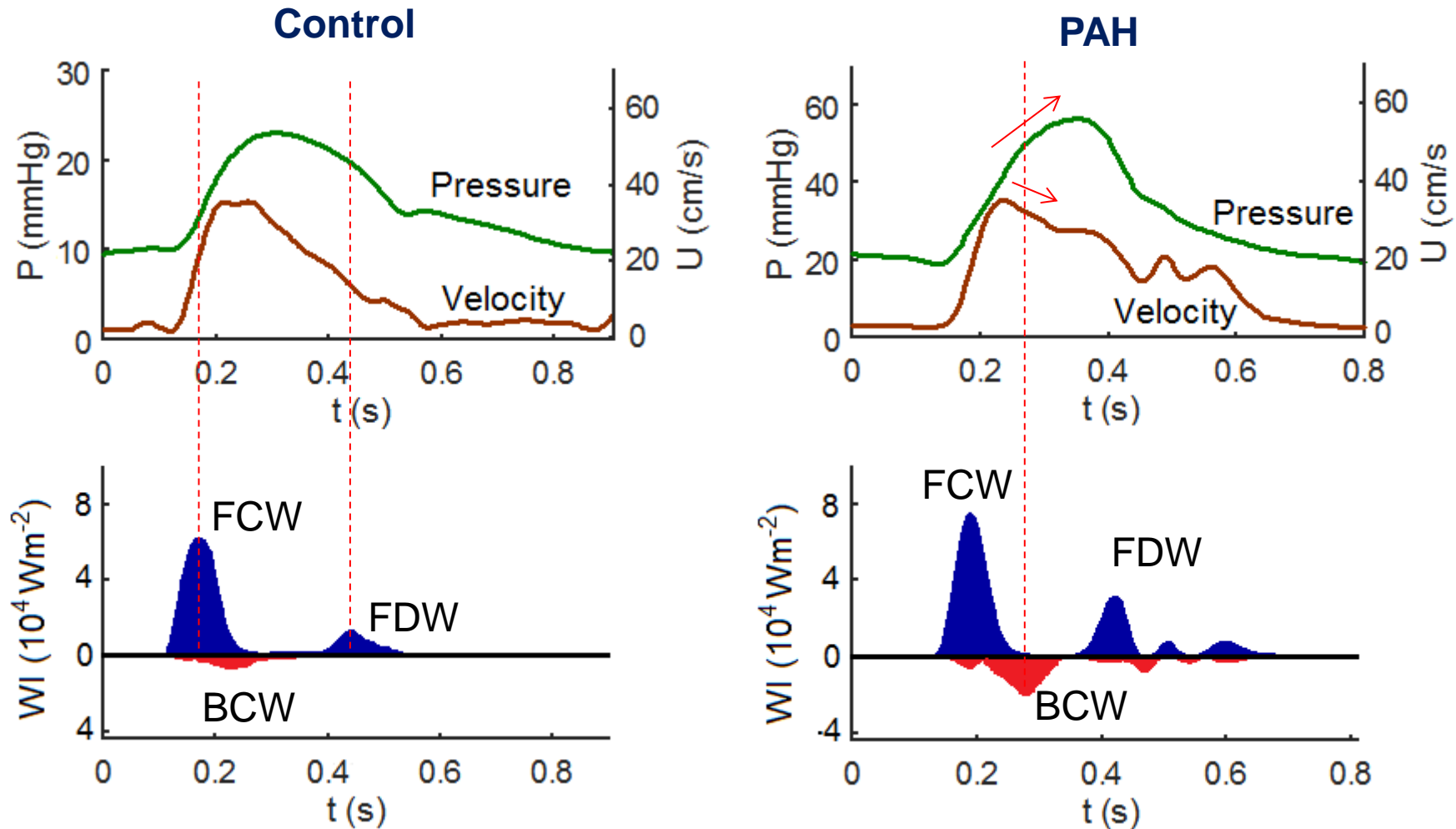


WIA pattern

Control

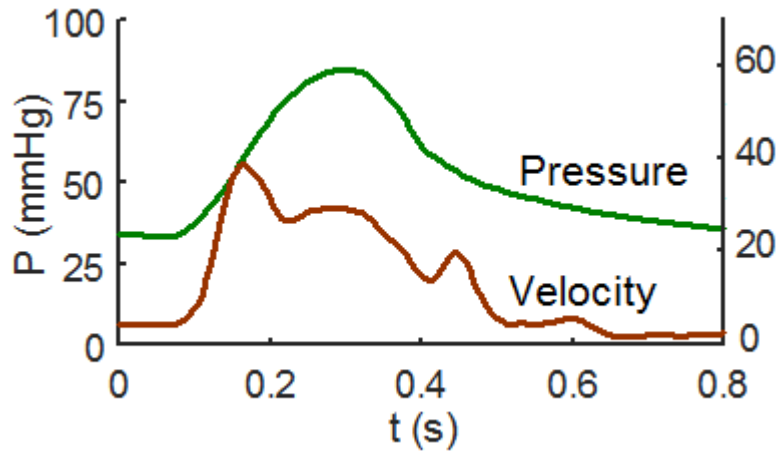


WIA pattern

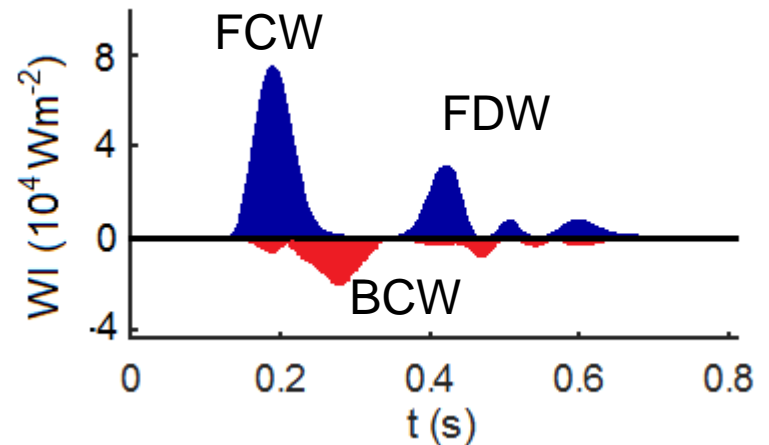
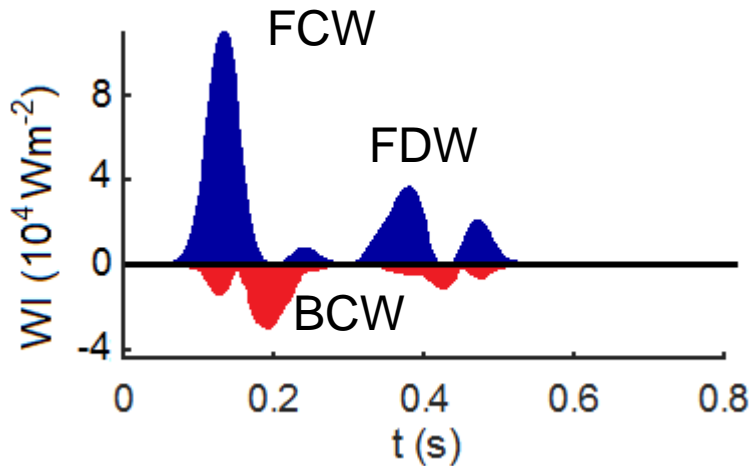
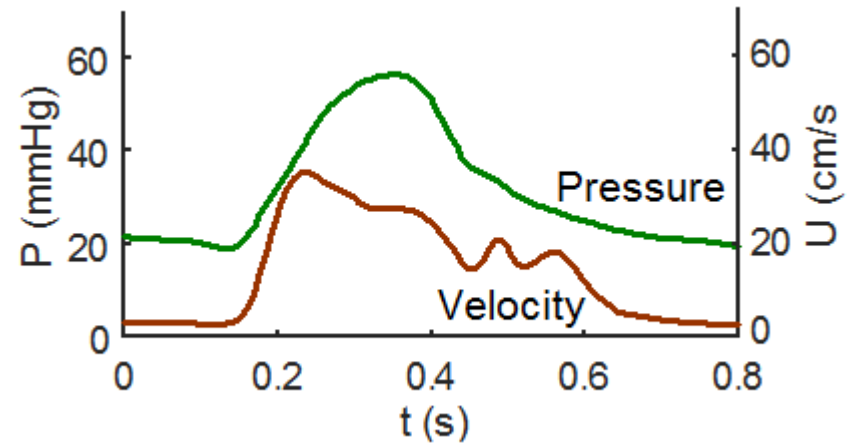


WIA pattern

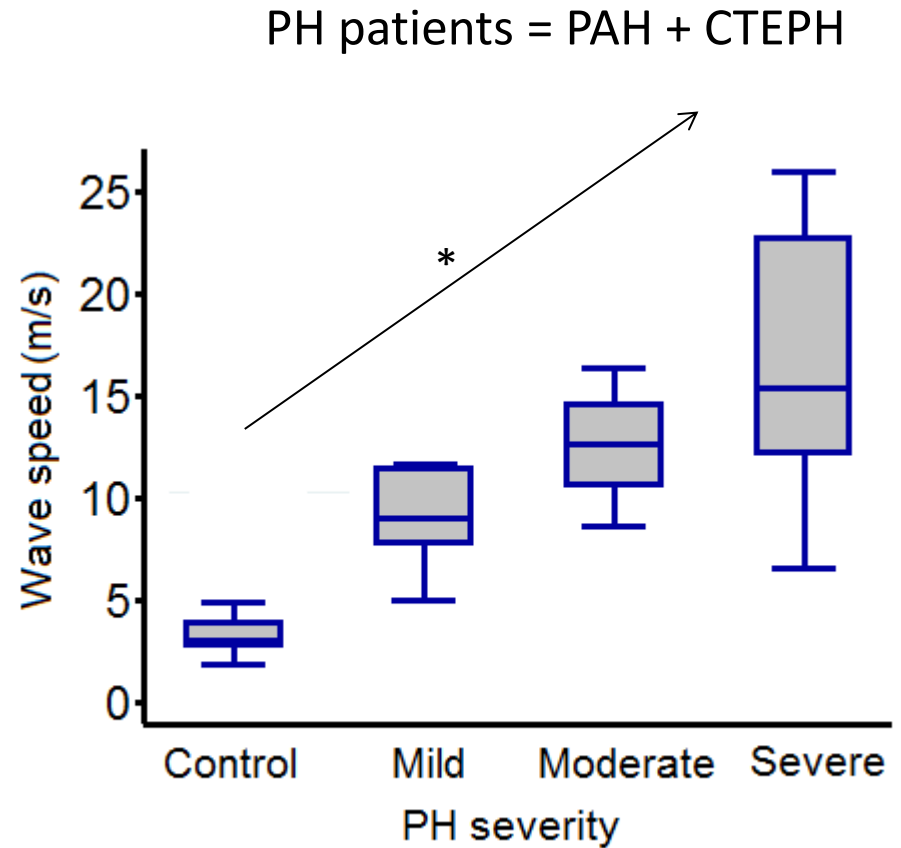
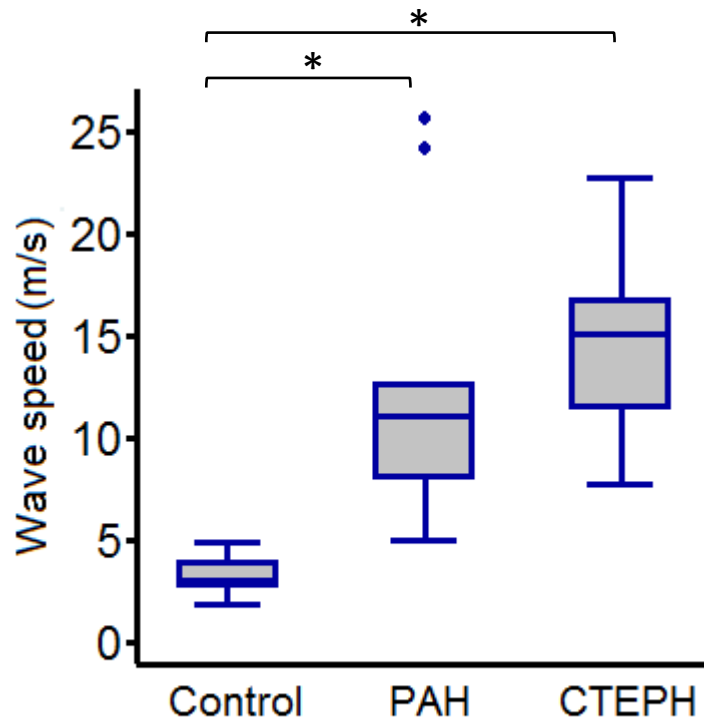
CTEPH



PAH



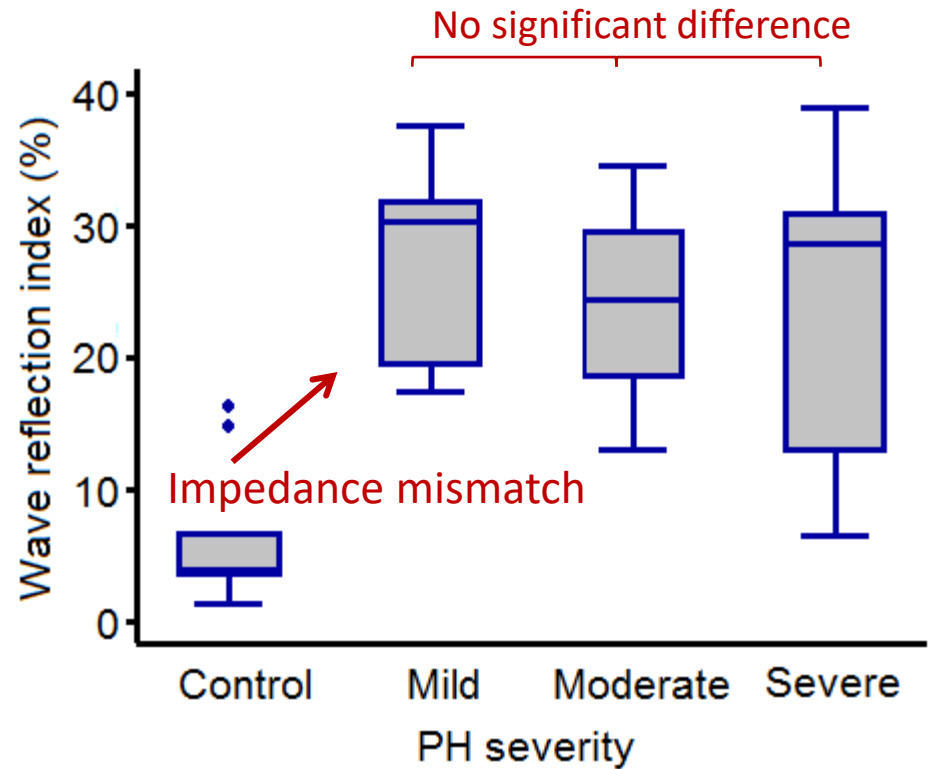
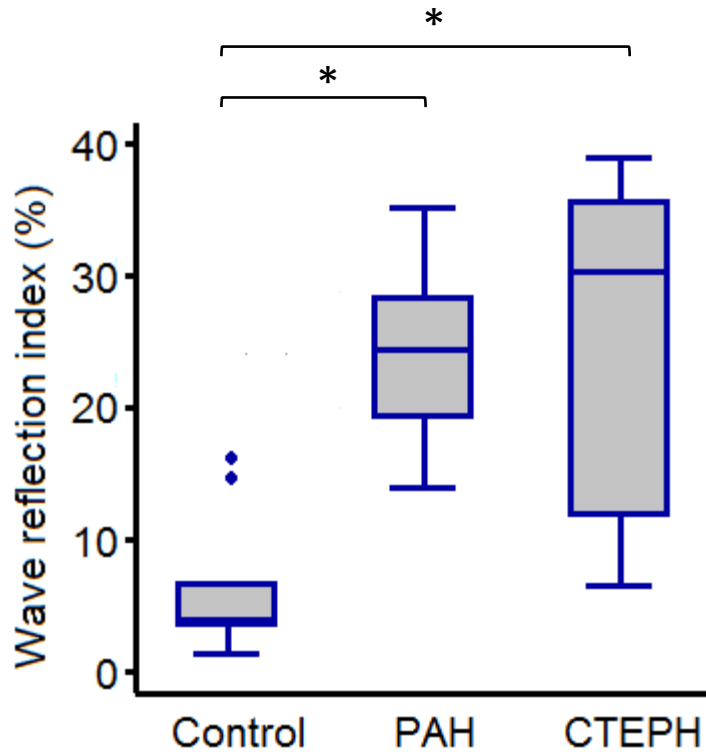
Wave speed



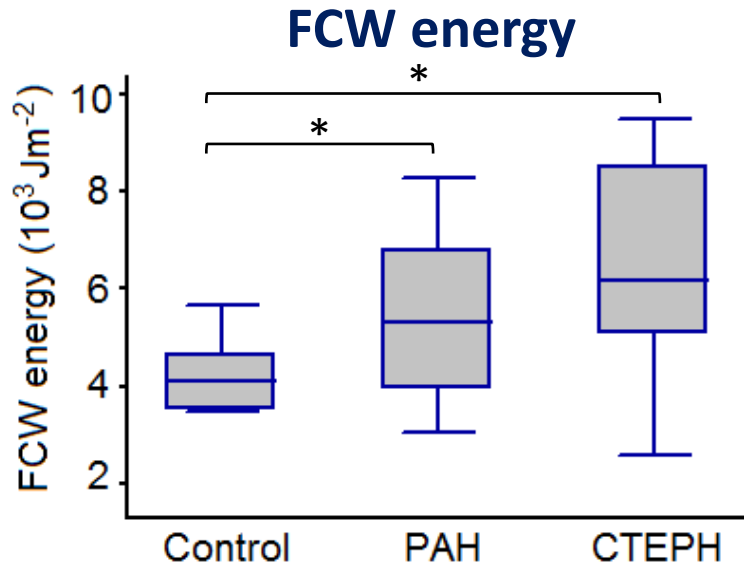
Wave reflection

Wave reflection index (WRI) = BCW/FCW

Wave reflection

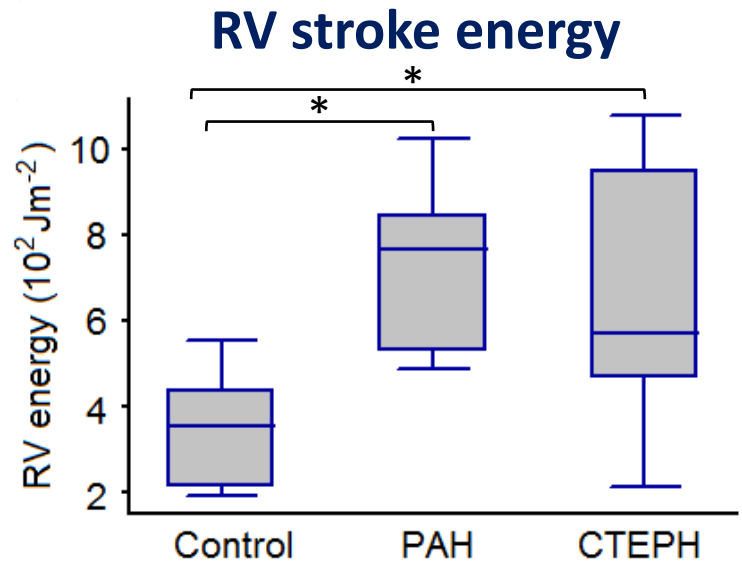
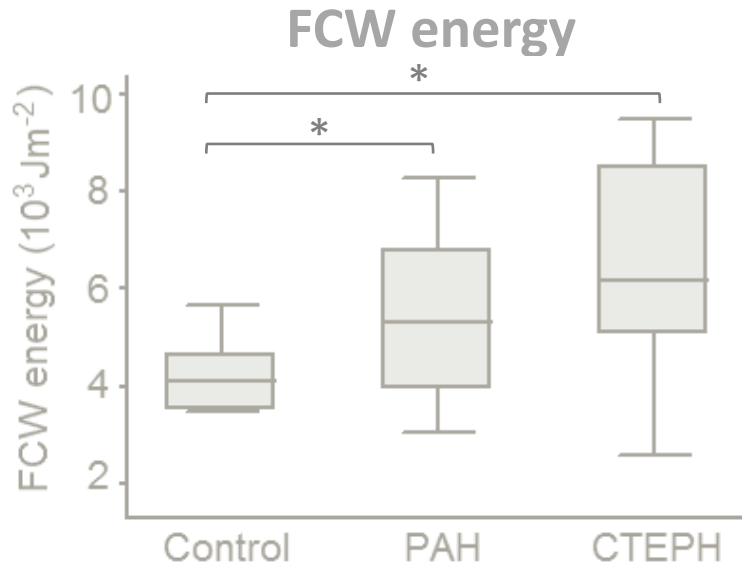


The right ventricle



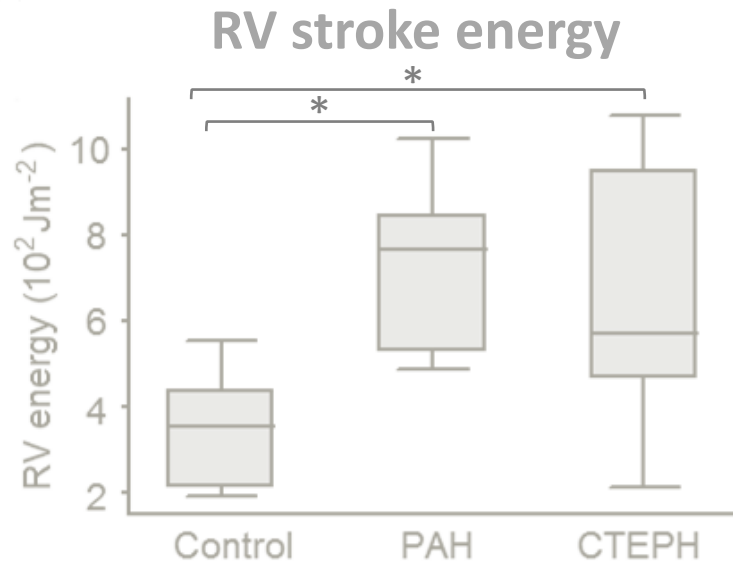
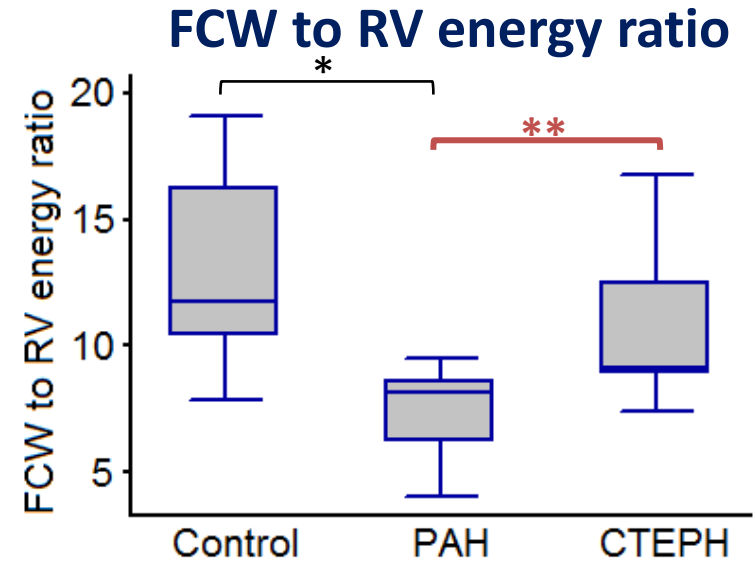
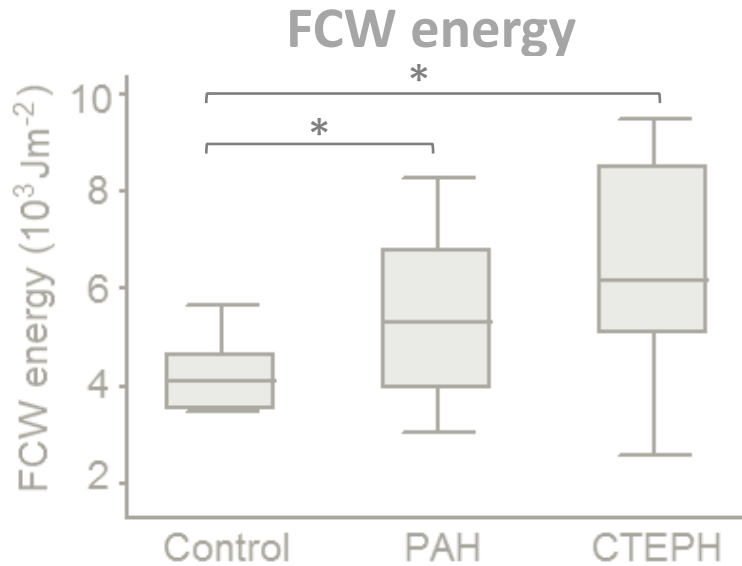
No significant difference

The right ventricle

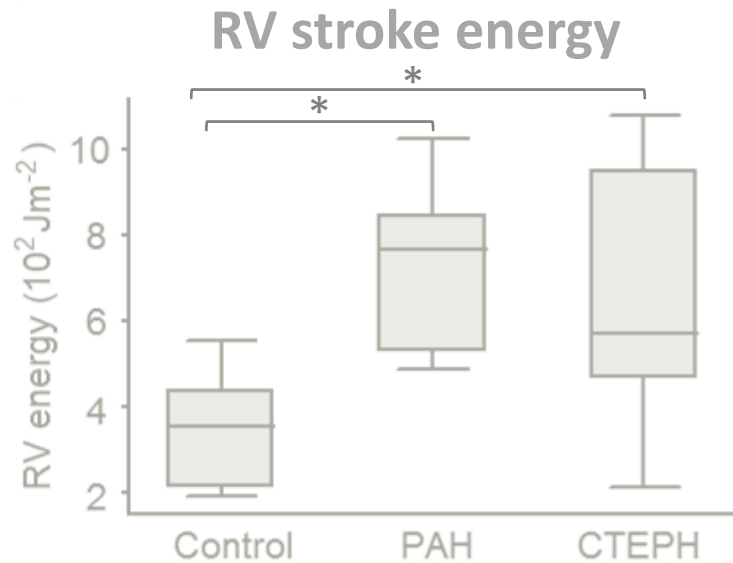
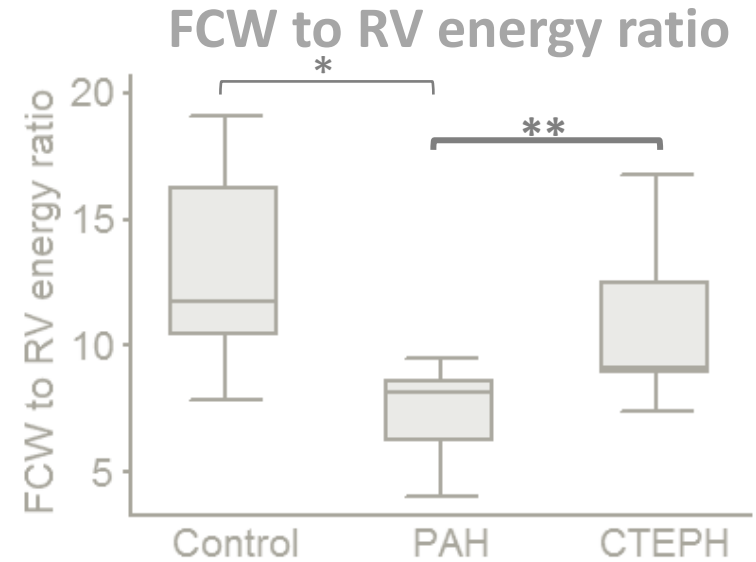
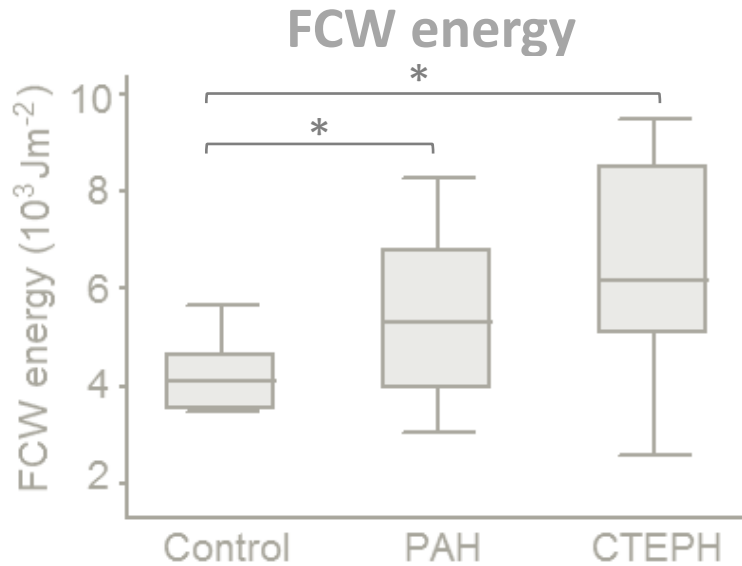


No significant difference

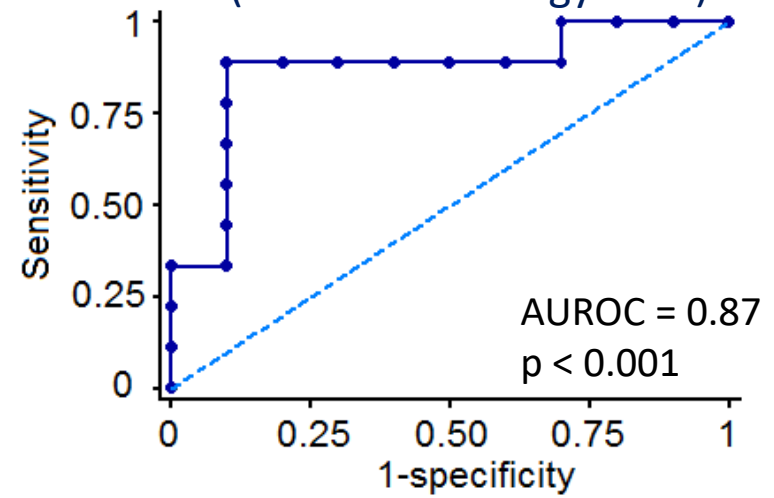
The right ventricle



The right ventricle



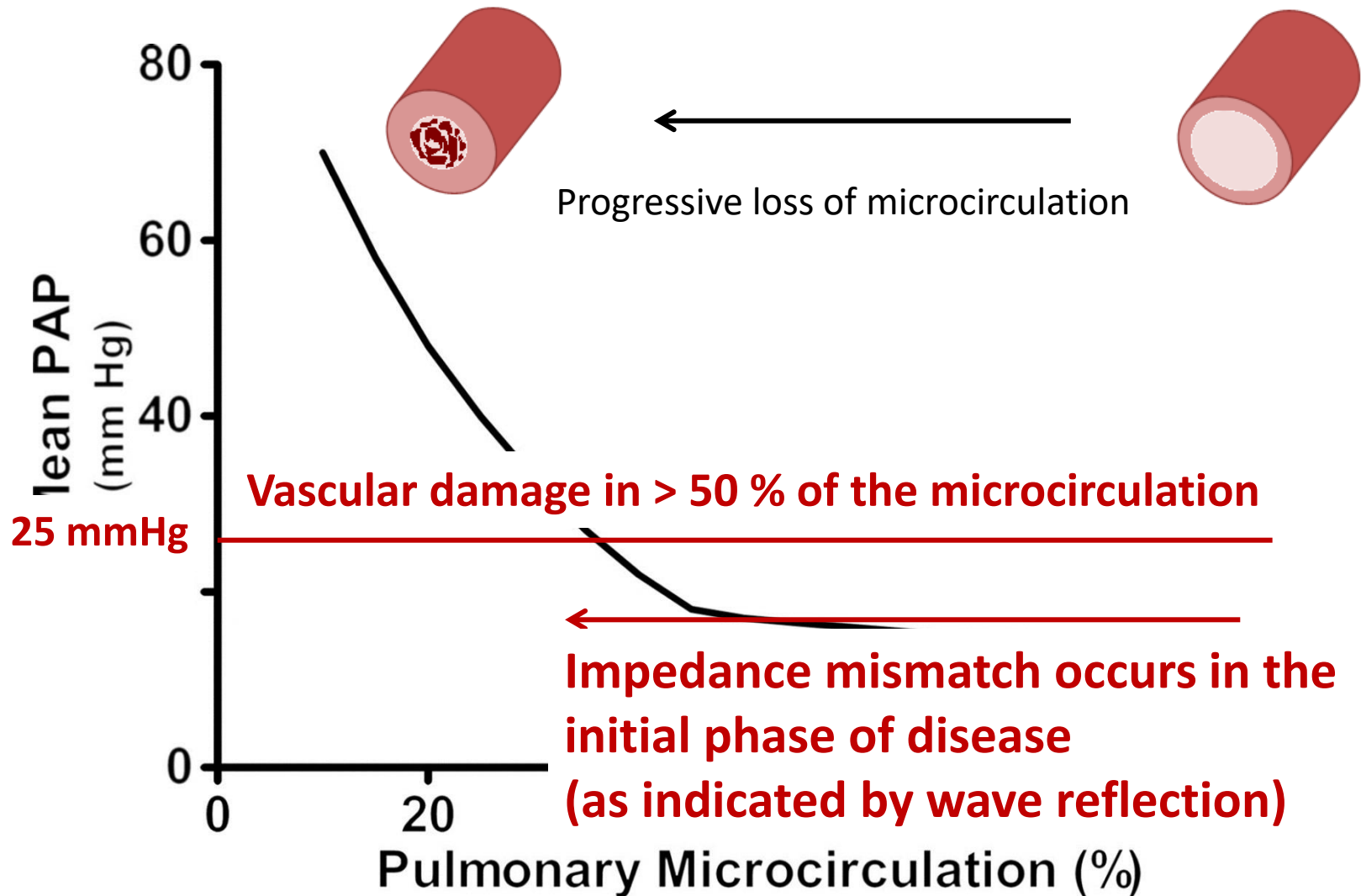
Differentiating CTEPH from PAH (FCW to RV energy ratio)





Discussion and conclusion

Early detection of disease



PAH versus CTEPH

- PAH: pharmacological treatment
- CTEPH: pulmonary endarterectomy



- FCW to RV energy ratio greater in CTEPH than PAH
 - ⇒ differences in RV function
 - CTEPH: rapid adaptation
 - PAH: gradual adaptation
- May serve as an additional measurement

Conclusion

- Wave speed increases in PH \Rightarrow greater arterial stiffness.
- Wave reflection is minimal in individuals without pulmonary vascular disease.
- Large wave reflection in pulmonary hypertension and it is unrelated to severity.
- Increased wave reflection may be an early indicator of pulmonary vascular disease.
- FCW to RV energy ratio can differentiate between PAH and CTEPH.



Thank you!

Acknowledgements

Aarhus group:

¹Prof Ulf Simonsen, ²Dr Jens Erik Nielsen-Kudsk, ³Dr Ole Hilberg,

²Dr Soren Mellemkjaer,

1. Department of Biomedicine, Aarhus University
2. Department of Cardiology, Aarhus University Hospital
3. Department of Respiratory Medicine, Aarhus University Hospital

London group:

^{1,2}Prof Alun Hughes, ³Prof Kim Parker, ¹Dr Charlotte Manisty, ²Dr

Luke Howard

1. Department of Cardiovascular Sciences, University College London
2. National Heart and Lung Institute, Imperial College London
3. Department of Bioengineering, Imperial College London

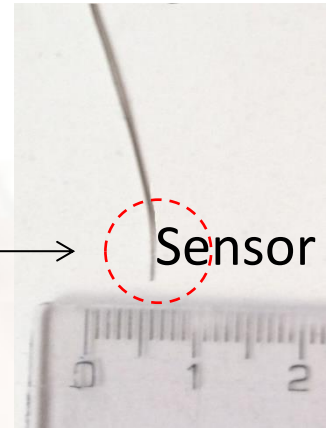
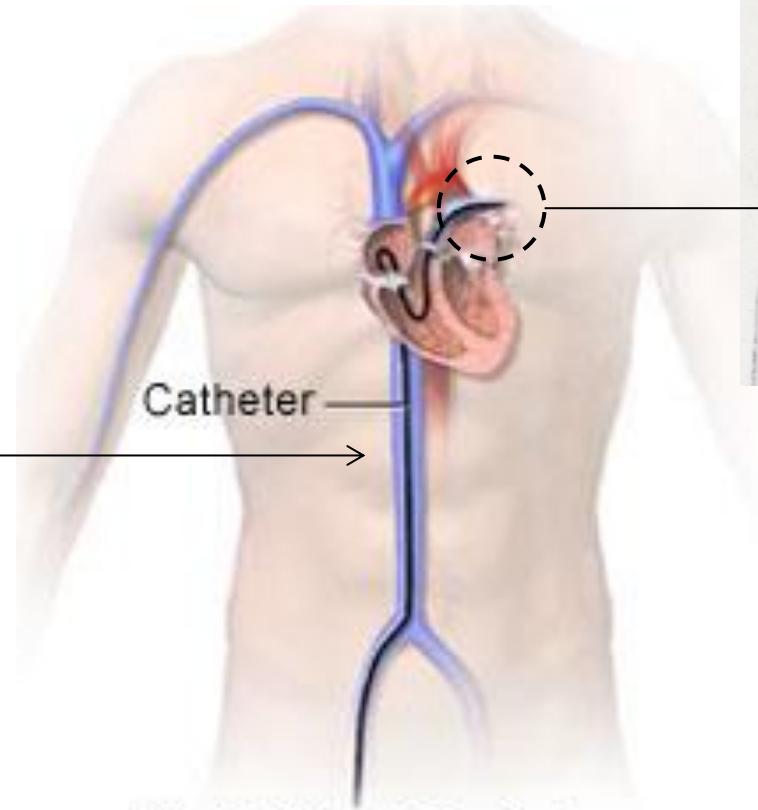
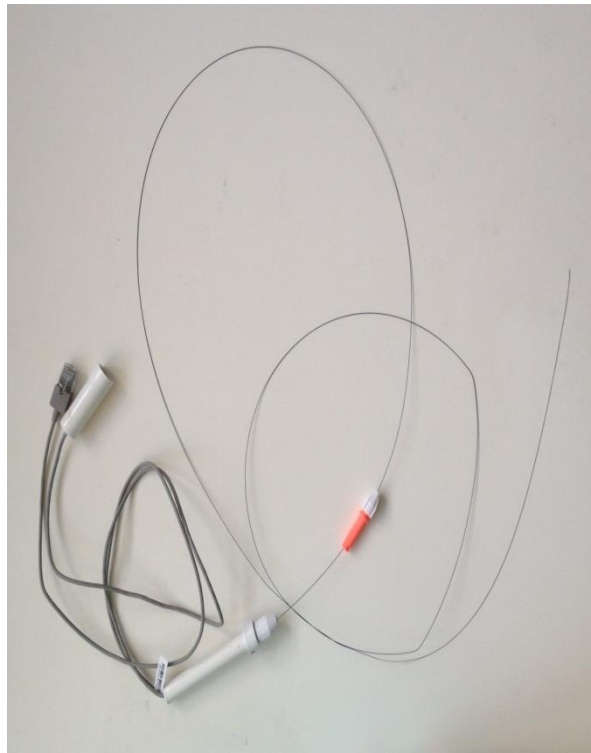
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Kornings Fond
Bønnelyckes Fond
Eva og Henry Frænkels Mindefond

Right heart catheterisation

Combowires



Right Heart Catheter

Calculations – wave intensity

“Original” wave intensity (Wm^{-2})

$$dI = dP \cdot dU$$

“Time-normalized” Wave intensity ($\text{Wm}^{-2}\text{s}^{-2}$)

$$WI = \frac{dP}{dt} \cdot \frac{dU}{dt}$$

Wave intensity normalized to sample numbers (Wm^{-2})

$$WI = dP \frac{CCD}{dt} \cdot dU \frac{CCD}{dt}$$

Calculations – RV energy

RV stroke work

$$RVS\!W = (PAP_m - RAP) \cdot RVS\!V$$

RV energy density

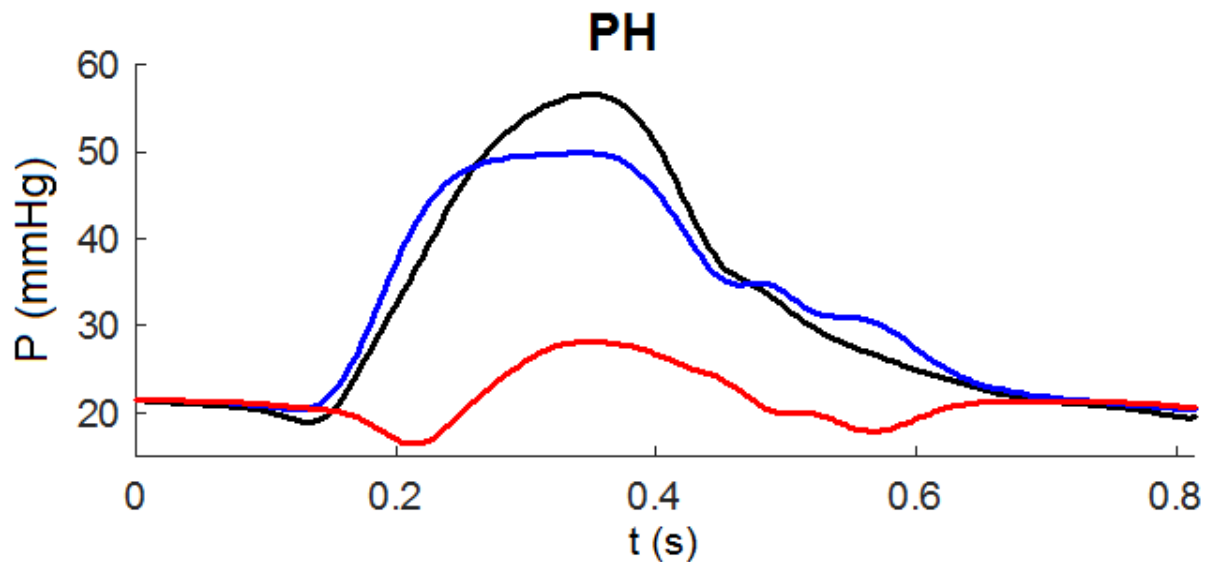
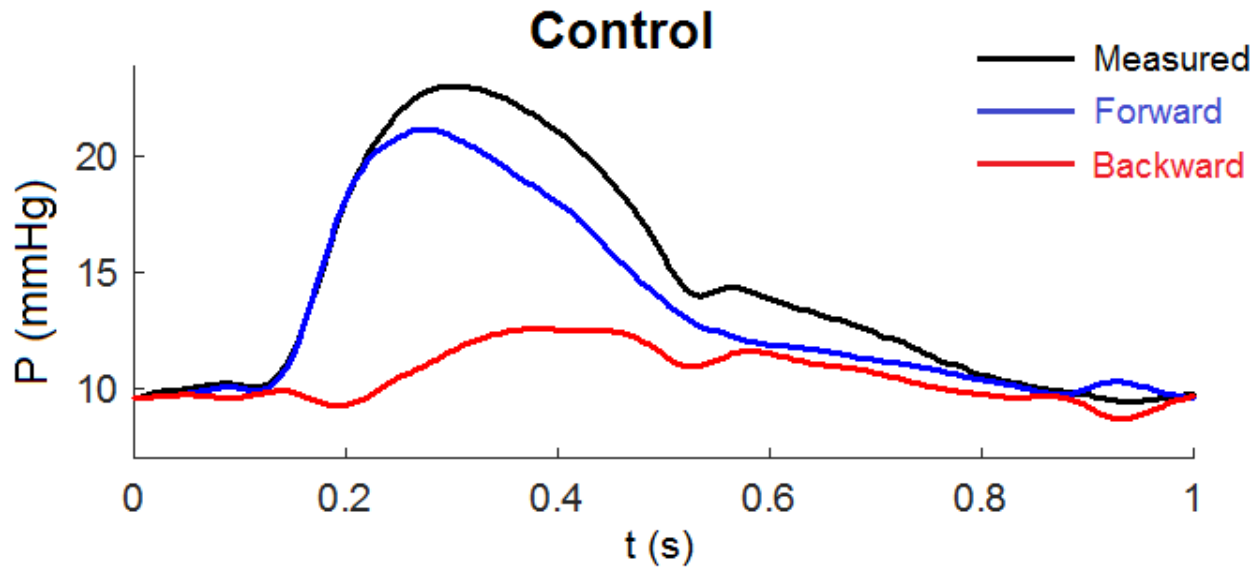
$$\begin{aligned} RV \text{ energy density} &= \frac{RVS\!W}{CSA} = \frac{(PAP_m - RAP) \cdot RVS\!V}{RVS\!V \cdot HR / U_{mean}} \\ &= \frac{(PAP_m - RAP)}{HR / U_{mean}} \\ &= (PAP_m - RAP) \cdot U_{mean} \cdot CCD \end{aligned}$$

Participant characteristics

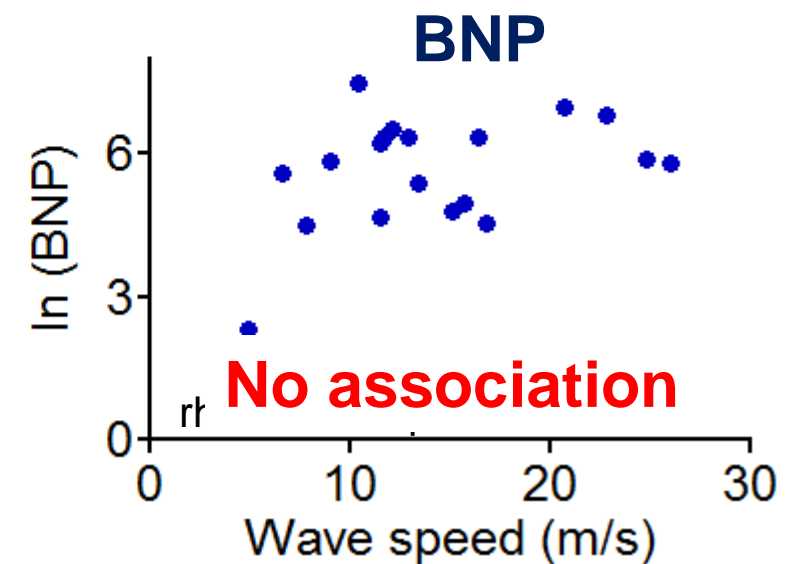
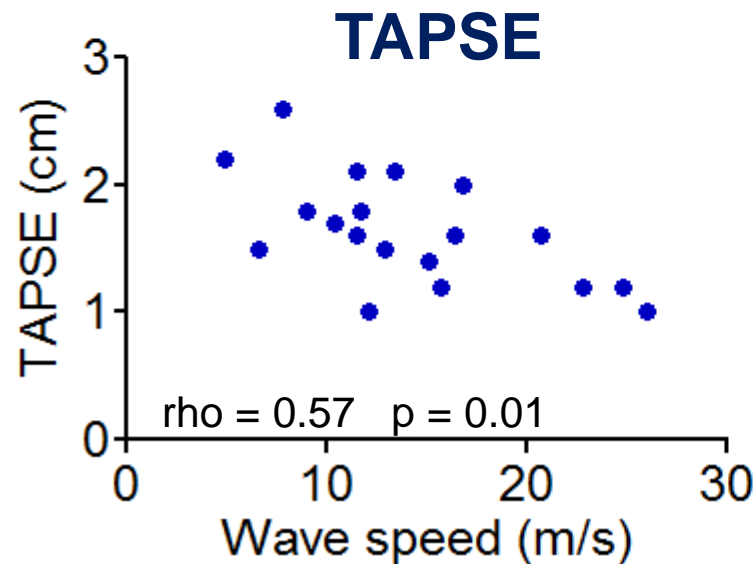
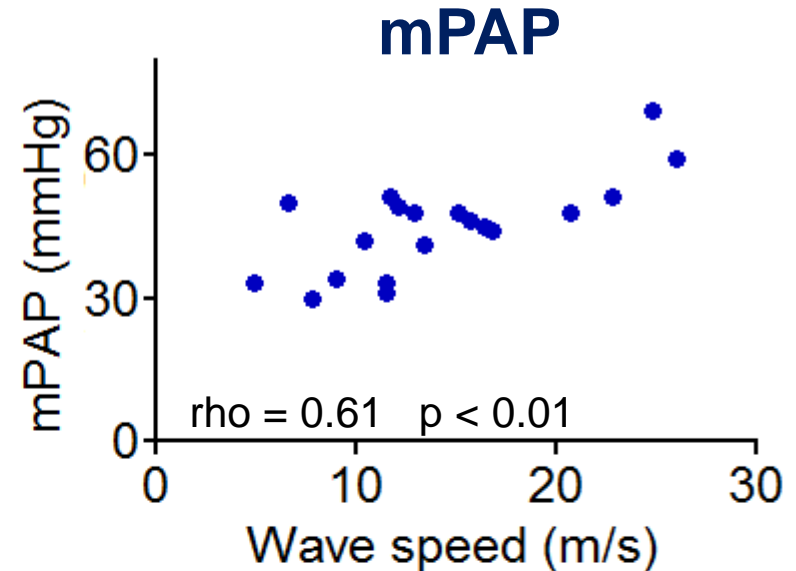
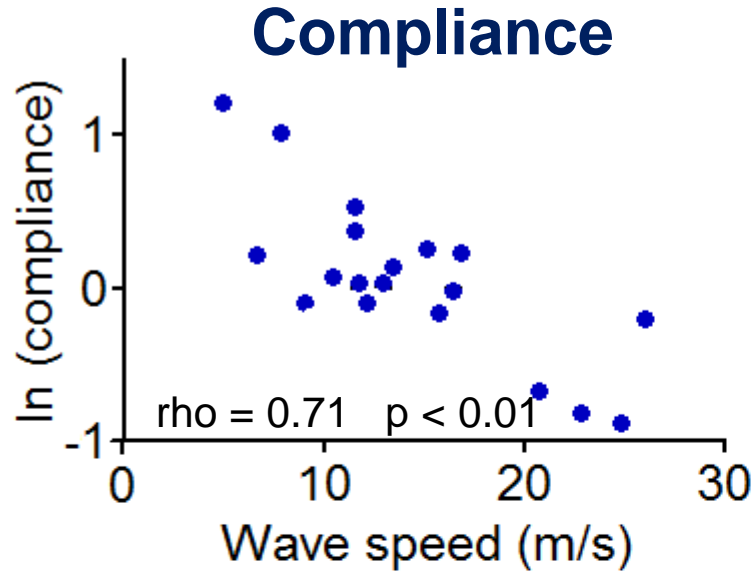
	Control N = 10	PAH N = 11	CTEPH N = 10
Age (yrs)	59 ± 14	56 ± 21	66 ± 9
Male, n (%)	8 (80 %)	2 (18 %)*	2 (20 %)*
BMI (kg/m ²)	28 ± 5	26 ± 5	27 ± 6
HR (beats/min)	73 ± 8	81 ± 8	80 ± 15
PAPm (mmHg)	17 ± 3	47 ± 11*	42 ± 8*
TPRI (WU/m2)	7 ± 2	25 ± 13*	20 ± 8*
CI, L/min/m2	2.6 ± 0.5	2.3 ± 1.1	2.4 ± 0.8
BNP (ng/L)	50 ± 64	522 ± 141*	265 ± 166*

*p < 0.05 vs control

Pressure separation



Wave speed



Wave reflection

