

X-RAY CT ASSESSMENT OF LUNG FUNCTION AND BIOMECHANICS

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Background

The lung is a complex, over-engineered organ comprising airways, blood vessels and parenchymal tissue. To ensure its main function of gas exchange, the lung constantly undergoes structural deformation through breathing and pulsatile blood flow. The biomechanical properties of viscoelastic respiratory tissues play a crucial role in both normal physiology and in diseases. For example, mechanical ventilation of patients with acute respiratory failure can cause excessive stress and strain and energy dissipation within the parenchyma leading to injury, which significantly contributes to mortality, a condition referred to as Ventilator-Induced Lung Injury (VILI)¹. The multiscale interaction of all the components that constitute the lung leads to a complex dynamic behaviour where lung function as a whole is not the sum of the behaviour of its individual components. Elucidating the lung microstructure and micromechanics *in vivo* in health and disease remains very challenging. X-ray CT is the state-of-the-art modality for imaging lung morphology. In the past decades, elastic image registration methods have been used to assess local lung biomechanics. However, clinical CT is limited in its spatial resolution. Synchrotron phase-contrast and K-edge subtraction CT offer unmatched capabilities in assessing local lung micromechanics and function, respectively. Although these techniques are not yet available in the clinical setting, they promise to further our understanding of lung biomechanics in experimental models.

Recent Advances

We previously introduced an energy-subtractive synchrotron CT technique that allows imaging regional lung function with unequalled spatial resolution *in vivo* in small animals². More recently, we have developed a synchrotron phase-contrast 4D- μ CT technique that allows the *in vivo* assessment of local lung strain under mechanical ventilation in both the parenchyma and blood vessels³ (Figure 1). In the clinical setting, work is underway to assess local lung biomechanical biomarkers in mechanically-ventilated patients with acute respiratory distress syndrome (ARDS) due to Covid-19, using advanced CT image processing methods, and to study their association with clinical outcome.

Future directions

Synchrotron phase-contrast 4D- μ CT is being used to assess lung micromechanics in disease models where it plays a critical role, such as VILI and fibrosis, where better understanding of the consequent lung

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extracellular matrix alterations and inflammation can offer new therapeutic and preventive opportunities. In the clinical arena, lung biomechanical parameters from patients with ARDS can be used to inform *in silico* lung models used to personalize mechanical ventilation settings for optimal lung protection, through “offline” simulation.

Figures

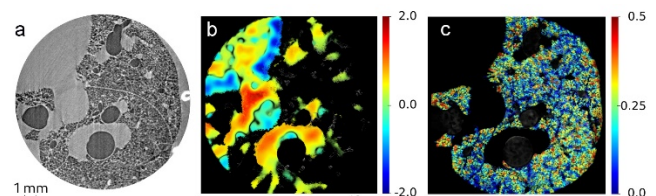


Figure 1: Quantitative mapping of lung tissue biomechanics in a live rat at $6 \mu\text{m}^3$ resolution. Sample X-ray phase-contrast CT image at 42 ms from start of inspiration (a); regional strain as a function of time computed within blood vessels (b) and airspaces (c). Modified from³.

References

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Acknowledgements

Alberto Bravin, Ludovic Broche, José Luis Cercos-Pita, Richard Deyhle, Luca Fardin, Irma Mahmutovic, Lars E. Olsson, Maciej Orkisz, Gaetano Perchiazzi, Jean-Christophe Richard, Mehdi Shekarnabi.

