## A phase I feasibility study in establishing the role of ultrasound-guided pleural biopsies in pleural infection (the audio study)

I Psallidas<sup>1</sup>, N Kanellakis<sup>1</sup>, R Bhatnagar<sup>2</sup>, R Ravindran<sup>1</sup>, A Yousuf<sup>1</sup>, AJ Edey<sup>3</sup>, RM Mercer<sup>1</sup>, JP Corcoran<sup>1</sup>, RJ Hallifax<sup>1</sup>, P Shetty<sup>1</sup>, T Dong<sup>4</sup>, HEG Piotrowska<sup>5</sup>, C Clelland<sup>6</sup>, NA Maskell<sup>2</sup>, NM Rahman<sup>1</sup>.

1 Oxford Centre for Respiratory Medicine, Churchill Hospital, Oxford University Hospitals NHS Foundation Trust, Oxford, UK

2 Academic Respiratory Unit, School of Clinical Sciences, University of Bristol, Bristol, UK

3 Department of Radiology, Southmead Hospital, North Bristol NHS Trust, Bristol, UK

4 Medical Research Council Human Immunology Unit, Medical Research Council Weatherall Institute of Molecular Medicine, Radcliffe Department of Medicine, Oxford, UK

5 Oxford Respiratory Trials Unit, Nuffield Department of Medicine, University of Oxford, Oxford, UK

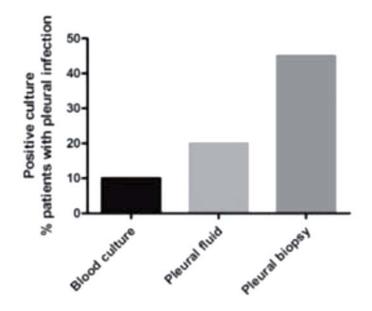
6 Department of Cellular Pathology, Oxford University Hospitals NHS Foundation Trust, Oxford, UK

**Background** Pleural infection is a common complication of pneumonia associated with high mortality and poor clinical outcome. Treatment of pleural infection relies on the use of broad-spectrum antibiotics, since reliable pathogen identification occurs infrequently. We performed a feasibility interventional clinical trial assessing the safety and significance of ultrasound (US)-guided pleural biopsy culture to increase the microbiological yield.

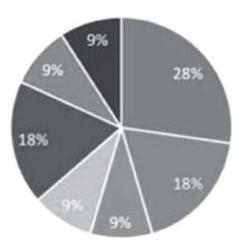
**Methods** 20 patients with clinically established pleural infection were recruited. Participants underwent a detailed US scan and US-guided pleural biopsies before chest drain insertion, alongside standard clinical management. Pleural biopsies and routine clinical samples (pleural fluid and blood) were submitted for microbiological analysis. In an exploratory sub-study, the 16S rRNA technique was applied on pleural biopsy samples, to investigate its' utility on increasing speed and accuracy versus standard microbiological diagnosis. This trials is registered with ClinicalTrials.gov, number NCT02608814

**Findings** US-guided were safe with no adverse events observed in this study. Pleural biopsies increased microbiological yield by 30% in addition to pleural fluid and blood samples (combined diagnostic sensitivity 55%). US characteristics at baseline were not statistically associated with survival, fluid volume drainage, radiological improvement or need for surgery. The 16S rRNA technique was successfully applied to pleural biopsy samples, demonstrating high sensitivity (93%) and specificity (89-5%).

**Conclusion** Our findings demonstrate safety of conducting US guided biopsies in patients with pleural infection and a substantial increase in microbiological diagnosis. qPCR primer assessment of pleural fluid and biopsy appears to have excellent sensitivity and specificity. Funding Oxfordshire Health Services Research Committee



% increase on positive culture samples in patients with pleural infection



- Streptococcus milleri
- Streptococcus intermedius
- Klebsiella pneumoniae

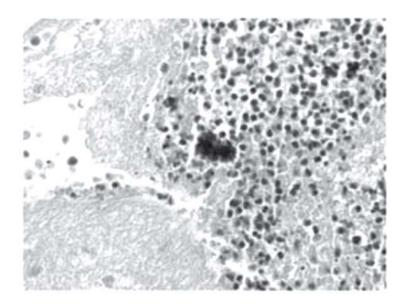
Staphylococcus aureus

Staphylococcus lugdunesis

Anaerobes

Staphylococcus Epidermidis

Results of pleural biopsy culture



Gram stain of acute inflammatory exudate in a pleural biopsy showing small colonies of Gram positive cocci.