

recommendations on dose. Recent studies have suggested that some critically ill patients fail to achieve sufficient plasma antibiotic concentrations to treat infection effectively.<sup>2</sup>

We determined whether critically-ill patients with respiratory infection achieved pharmacokinetic/pharmacodynamic (PK/PD) targets during antibiotic treatment and investigated factors associated with failure to meet these targets.

**Methods** This was a subgroup, interim analysis of an ongoing study, ABDose. Participants were adults in intensive care receiving piperacillin-tazobactam or co-amoxiclav for respiratory infection. Demographics and measures of organ function were recorded. Antibiotic concentrations were measured, at steady-state, in plasma at 50% and 100% of the dosing interval. Efficacy of beta-lactam antibiotics is dependent upon time above minimum inhibitory concentration (MIC). We chose PK/PD targets of antibiotic concentration >MIC and a more conservative >4 × MIC of likely pathogen or microbiological isolate (when available). These targets have been used previously.<sup>2</sup> During 28-day follow up, need for additional antibiotics was recorded.

**Results** 24 participants (median age 61, IQR [50–70] years), received co-amoxiclav (n = 7), piperacillin-tazobactam (n = 15) or both (n = 2). At 100% of the dosing interval, 12 achieved plasma antibiotic concentrations >MIC and 8 achieved >4×MIC. Participants who did not achieve PK/PD targets were younger (48 [39–59] years vs 68 [61–80] years, p = 0.002\*) and had a higher eGFR (131 ± 58 ml/min/1.73m<sup>2</sup> vs 64 ± 28 ml/min/1.73m<sup>2</sup>, p = 0.004\*) than those who did. Antibiotic concentrations were correlated with age and negatively correlated with eGFR (Figure 1). All participants failing to achieve antibiotic concentrations >4 × MIC at 100% of the dosing interval required further courses of antibiotics during follow-up compared to 50% of patients achieving this target (p = 0.02\*).

**Conclusion** In critically-ill patients with respiratory infection, uniform dosing of beta-lactam antibiotics does not consistently achieve PK/PD targets required for optimal efficacy. Younger patients, with better renal function may be under-dosed. These interim findings identify a need for further work to determine whether personalised dynamic dosing regimens could improve outcomes for patients with severe respiratory infection. Population PK modelling and further covariate analysis is planned within ABDose.

## REFERENCES

- 1 Mayr FB, et al. Epidemiology of severe sepsis. *Virulence* 2014;**5**:4–11.
- 2 Roberts JA, et al. DALI: defining antibiotic levels in intensive care unit patients: are current β-lactam antibiotic doses sufficient for critically ill patients?. *Clin Infect Dis* 2014;**58**:1072–83.

## S14 PATIENTS' PERCEPTIONS OF AN EXERCISE PROGRAMME DELIVERED FOLLOWING DISCHARGE FROM HOSPITAL AFTER CRITICAL ILLNESS (THE REVIVE TRIAL)

<sup>1</sup>K McDowell, <sup>2</sup>JM Bradley, <sup>3</sup>DF McAuley, <sup>2</sup>B Blackwood, <sup>1</sup>B O'Neill. <sup>1</sup>Centre for Health and Rehabilitation Technologies, Ulster University, Newtownabbey, UK; <sup>2</sup>School of Medicine, Dentistry and Biomedical Sciences, Queens University, Belfast, UK; <sup>3</sup>School of Medicine, Dentistry and Biomedical Sciences, Queens University and Regional Intensive Care Unit, Royal Victoria Hospital and Clinical Trials Unit, Belfast, UK

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**Introduction** The REVIVE RCT investigated the effectiveness of an individually tailored (personalised) exercise programme for patients discharged from hospital after critical illness.<sup>1</sup> By

including qualitative methods, we aimed to explore their perceptions of engaging in the 6 week programme to facilitate a better understanding of the intervention and trial outcomes.

**Methods** Patients allocated to the exercise group were invited to participate in semi-structured interviews following their final outcome assessment (6 months following randomisation). Interviews were conducted by a trained member of the research team not involved in the intervention. Interviews were audio recorded, transcribed verbatim and content analysis used to explore themes arising from the data.

**Results** Of 30 patients allocated to the exercise group 21 completed interviews. Seven core themes were identified (1) sequelae of critical illness and critical care recovery; (2) satisfaction and endorsement of the exercise programme; (3) beneficial impact of the exercise programme on physical and psychological health; (4) facilitators of beneficial impact; (5) barriers to beneficial impact; (6) challenges to continuing exercise; (7) contrasting views on outcome measures.

Patients provided insight into the physical and mental sequelae they experienced following critical illness. There was a strong sense of patients' need for the exercise programme and its importance for their recovery following discharge home. The programme was described as invaluable, and provided feelings of motivation and hope. Key facilitators of beneficial impact included supervision, tailoring of the exercises to personal needs, and the manual. Barriers to the beneficial impact of the programme included poor mental health, existing physical limitations and lack of motivation. Patients' views of the questionnaires and performance based outcome measures in the REVIVE trial varied. Many patients were unsure about what would be the best way of measuring how the programme affected their health.

**Conclusion** The benefits of physical rehabilitation programmes, needs to be counterbalanced against patients' mental health status post-ICU and any pre-admission limitations, if they are to be successful. Including this qualitative component improved our understanding of the mechanisms underpinning the impact of the programme and how programmes should be evolved for future trials.

## REFERENCE

- 1 O'Neill B, et al. Effectiveness of a programme of exercise on physical function in survivors of critical illness following discharge from the ICU: study protocol for a randomised controlled trial (REVIVE). *Trials* 2014;**15**:146.

## S15 CHANGES IN PERIOPERATIVE ARDS WITH TIME: A COMPARISON OF TWO TRIALS

<sup>1</sup>PA Howells, <sup>1</sup>KA Aldridge <sup>1</sup>RCA Dancer, <sup>2</sup>O Tucker, <sup>1</sup>DR Thickett. <sup>1</sup>Institute of Inflammation and Ageing, University of Birmingham, Birmingham, UK; <sup>2</sup>Department of Upper Gastrointestinal Surgery, Heart of England NHS Foundation Trust, Birmingham, UK

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**Introduction and objective** The BALTI-Prevention Trial1 translational sub-study (recruitment completed in 2011) and VINDALOO Trial2 (recruitment completed in 2015) both used oesophagectomy as a model for investigation of the pharmacological prevention of ARDS. The VINDALOO trial showed a lower ARDS incidence independent of the agents evaluated. Our objective was to characterise this difference.

**Methods** Databases from both trials were available and additional information was obtained retrospectively from hospital records. Analysis was performed using appropriate statistical tests.

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## S14 Patients' perceptions of an exercise programme delivered following discharge from hospital after critical illness (the revive trial)

K McDowell, JM Bradley, DF McAuley, B Blackwood and B O'Neill

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