**GODIF trial synopsis**

|  |  |
| --- | --- |
| **Title** | Goal directed fluid removal with furosemide in intensive care patients with fluid overload – A randomised, blinded, placebo-controlled trial (GODIF) |
| **Short title** | GODIF |
| **Objectives** | To assess the benefits and harms of fluid removal with furosemide vs placebo on patient-important outcome measures in adult ICU patients with fluid overload. |
| **Population** | Adult ICU patients with fluid overload of 5 % or more according to ideal body weight |
| **Interventions** | Furosemide infusion to achieve and maintain a neutral cumulative fluid balance. |
| **Comparator** | Placebo (saline 0.9%) |
| **Outcomes** | **Primary**:  Days alive and out of hospital at day 90 after randomisation.  **Secondary**:   1. All-cause mortality at day 90 after randomisation 2. Days alive at day 90 without life support (vasopressor/inotropic support, invasive mechanical ventilation or renal replacement therapy). 3. All-cause mortality at 1-year after randomisation 4. Number of participants with one or more serious adverse events (SAEs) and serious adverse reactions (SARs) to furosemide   **Exploratory**:   1. HRQoL 1-year after randomisation measured using the EuroQoL (EQ)-5D-5L and EQ-VAS scores. 2. Participants subjective assessment of their quality of life since the treatment in the ICU (unacceptable/neutral/acceptable) compared to (EQ)-5D-5L and EQ-VAS scores. 3. Cognitive function 1-year after randomisation as assessed by the Montreal Cognitive Assessment (MoCA mini) score. |
| **Eligibility** | **Inclusion criteria:**   1. Acute admission to the ICU **AND** 2. Age ≥ 18 years of age **AND** 3. Fluid accumulation ≥ 5% of ideal bodyweight *estimated* according to the daily fluid charts, the cumulative fluid balance, development in body weight, and clinical examination **AND** 4. Clinical stable assessed by clinicians (minimum criteria: MAP > 50 mmHg and maximum infusion of 0.20 microgram/kg/minute of noradrenaline and lactate < 4.0 mmol/L).   **Exclusion criteria:**   1. Known allergy to furosemide or sulphonamides 2. Known pre-hospitalization advanced chronic kidney disease (eGFR < 30 mL/minute/1.73 m2 or chronic RRT) 3. Ongoing RRT 4. Anuria for ≥ 6 hours 5. Rhabdomyolysis with indication for forced diuresis 6. Ongoing life-threatening bleeding 7. Acute burn injury of more than 10 % of the body surface 8. Severe dysnatremia (p-Na < 120 or > 155 mmol/l) 9. Severe hepatic failure as per the clinical team 10. Patients undergoing forced treatment 11. Pregnant women 12. Informed consent not obtainable |
| **Sample size** | 2 x 500 (15 % relative risk reduction (8 % absolute) in furosemide vs. placebo group in the primary outcome, assuming a baseline 90-day mortality of 27 % (two sided α=0.05 and β=0.2) |
| **Trial duration** | The trial intervention will continue for a maximum of 90 days post-randomisation. 90 days and 1-year follow-up post randomisation. The trial was initiated August 2020. |