

26 February 2024
Clinical Trials Unit Update

1. Status – Public**2. Executive Summary**

This paper provides an update on the clinical trials in organ donation and transplantation that NHSBT Clinical Trials Unit (CTU) are currently managing. There are currently 8 trials, which are described in the table below.

Key highlights since previous RINTAG report:

- The protocol for the HOPP study (Hypothermic Oxygenated Pancreas Perfusion) is undergoing Sponsor review and is currently aiming to begin recruitment in June 2024. The protocol will be shared for review by ROFG upon Sponsor approval.
- The DeFat study opened to recruitment on 23/02/2023, all 5 sites are currently open and 15 livers have been randomised to the study.
- The SIGNET study is continuing to open sites to recruitment. All of the planned 92 Trusts are now open, with 1,026 organ donors randomised so far.
- the COBALT study has completed recruitment (n=40). The NIHR have approved extension of the project until 31st March 2024 for close-down and analysis.
- The results from the PLUTO trial have now been published in Kidney International <https://doi.org/10.1016/j.kint.2023.09.032>
- The PITHIA analyses have been performed and are being considered for peer-review publication.

3. Action Requested

The Committee is asked to:

- Note the contents

Table 1. Current OTDT studies managed by the CTU
(in **set-up**, **recruitment**, or **follow-up/analysis**)

Trial Name	Trial Summary	Chief Investigator(s)	Sponsor	Progress
HOPP	Investigate the feasibility and safety of Oxygenated Hypothermic Machine Perfusion of pancreas grafts prior to transplantation in clinical practice.	Prof James Hunter	University of Oxford	Protocol and other key documents are currently undergoing review by Sponsor. Will be submitted to ROFG once Sponsor approval provided.
SIGNET-EME	Examine the mechanisms in the potential down regulation of brain stem death induced proinflammatory responses by Simvastatin	Prof Simi Ali	The Newcastle upon Tyne Hospitals NHS Foundation Trust	A mechanistic sub-study of SIGNET, using QUOD samples. In early set-up stage.
DeFat	Delivery of Ex-situ deFattening Agents during normothermic liver perfusion for Transplantation– a randomised clinical trial.	Prof Peter Friend Mr Simon Knight Mr Syed Hussain Abbas	University of Oxford	Open to recruitment since February 2023. To date, 5 sites are open to recruitment and 15 livers have been enrolled to the study. The NIHR have just approved a 20-month extension, therefore new end date of November 2025.
SIGNET	Statins for Improving orGaN outcomE in Transplantation.	Prof John Dark Prof Dan Harvey	The Newcastle upon Tyne Hospitals NHS Foundation Trust	SIGNET opened to recruitment in September 2021. To date, 92 Trusts have opened to recruitment, and 1,026 donors have been recruited to date.
TWIST	Multicentre, Open-label Randomised Trial of Superficial Wound Drain on Surgical Site Infection in high Body Mass Index (BMI) Kidney Transplant Recipients.	Mr James Hunter	Oxford University Hospitals NHS Foundation Trust	Providing administrative support to this study. There are currently 11 centres open to recruitment and recruitment continues.
COBALT	Cardiorespiratory Optimisation By AVF Ligation after Transplantation.	Prof Gavin Pettigrew	Cambridge University Hospitals NHS Foundation Trust & the University of Cambridge	The 40 th and final participant was recruited in April. Due to the extended recruitment period, a contract extension up to 31 st March 2024 has been approved by the NIHR.
PLUS	Utilisation of normothermic machine preservation in extended criteria livers – a national threshold-crossing study.	Prof Peter Friend Mr Simon Knight	University of Oxford	All 7 UK liver transplant centres are participated in the study. The recruitment target of 1035 livers was reached in April 2023. Follow up is ongoing until May 2024
PITHIA	Pre-Implantation trial of Histopathology In renal Allografts	Prof Gavin Pettigrew	Cambridge University Hospitals NHS Foundation Trust & the University of Cambridge	The analysis has been performed and results presented to the Investigators and PPI representatives in December 2023. The NIHR report will be submitted on 28 th Feb and the manuscript is in draft.

Author
CTU team
February 2024