

13th December 2022

Dear Research Ethics Committee,

Trial Title: Statins for Improving orGaN outcomE in Transplantation**Acronym: SIGNET****CI's: Professor John Dark and Dr Dan Harvey****Sponsor: The Newcastle Upon Tyne Hospitals NHS Foundation Trust****Sponsor ref: 09691****IRAS ref: 288722**

We are pleased to enclose Substantial Amendment 01 for the SIGNET trial for REC review.

This amendment to the protocol addresses the following revisions:

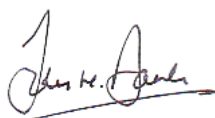
1. Inclusion of level 3 Trusts.
2. Clarification that consent for SIGNET may be taken prior to neurological death testing if this is deemed appropriate by the consenting SNOD, but donors will not be randomised to the study until after confirmation of neurological death. This follows feedback from the Specialist Nurses for Organ Donation and a review of eligible donors not randomised to SIGNET where we found that some donor families will consent for organ donation prior to neurological testing and then request no further contact.
3. Changes to the statistics section to clarify what will happen for missing registry data and the analysis population.
4. Clarification of source data for SIGNET.
5. Administrative changes.

Please find the following submitted documents for your review.

Document	Date	Version
09691 SIGNET Protocol Tracked Changes	07/12/2022	2.0
09691 SIGNET Protocol Clean	07/12/2022	2.0
09691 SIGNET Amendment Tool SA01 Protocol Change	13/12/2022	1.0

We would be most grateful for the REC's consideration and review of this amendment. Please do not hesitate to contact us if you have any queries.

With Kind Regards

Prof John Dark
Co-Chief Investigators (SIGNET Study)

Dr Dan Harvey