

05/08/2026

IRAS Reference: 1008743

Study Title: Precision medicine Adaptive Network platform Trial in Hypoxemic acute respiratory failure, the PANTHER Trial

Protocol number: 175151

Protocol Version: v2.1

Dear REC and MHRA,

PANTHER Substantial Modification 02 – SM02

We would like to submit a substantial modification to make the following changes:-

1. Master protocol updates - main changes listed below:-

- Clarification of blood gas parameters in the eligibility assessment of ARDS
- Clarifications around safety outcome reporting (within 28 days in line with SAEs)
- Addition of New-Onset Atrial Fibrillation (NOAF) as a safety outcome
- ECMO added as primary endpoint
- Addition of ventilator support as a secondary endpoint
- Clarification that patients will receive usual care or usual care + intervention
- Update to protocol to confirm medians for reporting
- The addition of the Brief Health Literacy Screening Tool, Labour market outcomes- including return to work, time off work and reason for time off work, as tertiary outcome

2. Update to original application to reflect that PANTHER Is running in the US, Japan, EEA, Canada & Australia

3. Request to extend labelling exemption originally granted for the study under the new Clinical Trial Regulations.

4. Correction, the impact assessment scale (6pt scale) is being submitted in replacement of the previously submitted 22 point scale.

- Addition of the IQCODE questionnaire
- Addition of the Brief Health Literacy Screening Tool

5. Update to UK Region Specific Appendix (RSA) to change the approval of the UK RSA (and all subsequent country RSAs) from the ITMG to the Executive Committee.

6. Updated SmPCs submitted

The below documents are included in this submission:

- PANTHER Master Protocol Summary of Changes v1.0 22.07.2026 SM02
- PANTHER Master Protocol V3.0 22.07.2026_Tracked
- PANTHER Master Protocol V3.0 22.07.2026_Clean
- PANTHER_UK_Region Specific Appendix_v2.0 01.06.2026 TC
- PANTHER_UK_Region Specific Appendix_v2.0 01.06.2026 Clean
- PANTHER SM02 Modification_Tool_v2_1_24Apr26
- Brief Health Literacy Screening Tool v1.0 10.10.2023
- PANTHER IQCODE v1.0 22.07.2026
- Impact of events scale 6pt v1.0 22.07.2026
- PANTHER Annex 13 Labelling Exemption Request v1.0 26.06.2026
- Simvastatin SmPC_updated 12.06.2026
- Baricitinib_SmPC_(Olumiant)_updated 03.06.2026
- PANTHER Summary of SmPC Changes v1.0 22.07.2026 SM02

 Digitally signed by Janis
Best-Lane
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