

The enhanced antenatal care bundle to improve maternal and newborn outcomes in Malawi and Zambia: a multi-country, parallel cluster-randomized trial

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Registration date 13/03/2025	Overall study status Ongoing	<input type="checkbox"/> Statistical analysis plan <input type="checkbox"/> Results
Last Edited 08/07/2025	Condition category Pregnancy and Childbirth	<input type="checkbox"/> Individual participant data <input checked="" type="checkbox"/> Record updated in last year

Plain English summary of protocol

Background and study aims

Even though many pregnant women in Malawi and Zambia receive regular healthcare during pregnancy and give birth in health centres, there are still many women who die from complications during childbirth. We have reliable evidence that these maternal deaths can be prevented with good quality healthcare that follows the guidance of the World Health Organisation (WHO). The WHO recommend that all pregnant women have regular health visits during pregnancy; the WHO also recommends that healthcare staff perform specific checks during each pregnancy visit so that any problems during pregnancy are found and treated before they become life-threatening to women or their babies.

This study will look at the healthcare pregnant women receive during their pregnancy and when they give birth in primary health centres in Malawi and Zambia. The researchers will compare important events during pregnancy and birth in facilities where staff provide women with standard pregnancy care versus those same important events in facilities where staff deliver an improved pregnancy healthcare service. The improved pregnancy health services include training for staff and pregnant women and supplies for health facilities.

Who can participate?

This study will include health facilities rather than individuals. The health centres will have 700 - 2000 visits per year from pregnant women. The health centres will also be able to provide emergency pregnancy care, provide services for free to pregnant women, and complete an assessment process to ensure that the site is ready for the trial.

What does the study involve?

The study will start with a baseline phase, where the researchers will measure key events during pregnancy and childbirth. Next, the facilities will be randomly assigned to either continue providing their standard pregnancy care with basic supplies or to deliver the study intervention to improve pregnancy care, with training for staff and pregnant women and with basic supplies.

What are the possible benefits of participating?

The researchers expect that the health centres receiving the study intervention to improve pregnancy will be better able to provide care that meets WHO standards. They expect these health facilities to have better results for women and babies in terms of avoiding complications during pregnancy that lead to preventable deaths amongst women.

Where is the study run from?

This study is run from the Malawi Liverpool Wellcome Institute in Blantyre (Malawi) and from Lusaka Apex Medical University in Lusaka (Zambia). The study teams also have support from the University of Liverpool (UK).

When is the study starting and how long will it run for?

August 2022 to July 2026

Who is funding the study?

National Institute for Health and Social Care Research (UK)

Who is the main contact?

safemotherhood@liverpool.ac.uk

Contact information

Type(s)

Principal Investigator

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Additional identifiers

EudraCT/CTIS number

Nil known

IRAS number**ClinicalTrials.gov number**

Nil known

Secondary identifying numbers

University of Liverpool Sponsor UoL001883

Study information

Scientific Title

The enhanced antenatal care bundle to improve maternal and newborn outcomes in pregnant women attending maternity health services in Malawi and Zambia: a multi-country, parallel cluster-randomised trial

Acronym

ENHANCE

Study objectives

ENHANCE will provide a comprehensive bundle of measures carefully developed for primary health centres in resource-limited settings, to support the consistent and comprehensive

delivery of all aspects of antenatal care according to WHO guidelines. ENHANCE includes tools, leadership, mentorship, task shifting and basic antenatal care supplies.

This study will determine whether an ENHANCE strategy compared with standard antenatal care is acceptable to pregnant women and healthcare staff; whether ENHANCE improves the implementation of antenatal care services according to WHO guidance; and whether ENHANCE improves both the quality of antenatal care and the maternal and neonatal clinical outcomes for women and babies who receive the intervention.

Ethics approval required

Ethics approval required

Ethics approval(s)

1. Approved 24/02/2025, The University of Liverpool Ethics Committee (Foundation Building, Liverpool, L69 7ZX, United Kingdom; +44 (0)151 794 2000; research.ethics@liverpool.ac.uk), ref: 15682

2. Approved 05/12/2024, College of Medicine Research Ethics Committee (Private Bag 360 Chichiri, Blantyre, 3, Malawi; +265 (0)1874377; comrec@medcol.mw), ref: P.10/24-1155

3. Approved 25/10/2024, Lusaka Apex Medical University Biomedical Research Ethics Committee (Foxdale Campus Plot No. 3073, Lusaka, PO Box 31909, Zambia; +260 (0)979452488; lamubrec@lamu.edu.zm), ref: 0294/25/10/2024

4. Approved 23/01/2025, National Health Research Authority (Lot No. 18961/M off Kasama Rd, Chalala Lusaka, PO Box 30075, Zambia; +260 (0)211250309; znhrasec@nhra.org.zm), ref: NHRA-1849/09/01/2025

Study design

Parallel cluster-randomized trial

Primary study design

Interventional

Secondary study design

Cluster randomised trial

Study setting(s)

GP practice, Hospital, Medical and other records

Study type(s)

Prevention, Treatment

Participant information sheet

Not available in web format. Please use contact details to request a participant information sheet.

Health condition(s) or problem(s) studied

Delivery of antenatal care services

Interventions

The study will implement a package designed to improve the quality and content of antenatal care delivered in resource-limited settings and compare this with the standard antenatal care offered to pregnant women in Malawi and Zambia.

This is a multi-country, parallel cluster-randomised controlled trial with a baseline period. There is an integrated implementation evaluation and health economic evaluation. The trial will include primary healthcare centres in Malawi and Zambia as clusters. During the first 3 months each cluster will provide standard care to establish weekly rates of maternal and newborn outcomes. After the baseline, the clusters will be randomly allocated in a 1:1 ratio to ENHANCE or standard care with the provision of basic ANC supplies, using a minimisation algorithm to account for the imbalance in location, hospital deliveries volume and distance to referral facility. Both quantitative and qualitative approaches will gather the data required to meet the objectives. The researchers will conduct an intention-to-treat analysis of the composite primary outcome. Descriptive statistics for continuous (mean and standard deviations), discrete (median and range or interquartile range) and categorical (frequency counts and percentages) variables will summarize baseline characteristics (demographic and clinical data) and outcome variables. Process evaluation surveys will be summarised using descriptive statistics as appropriate, and responses compared across PHCs and countries. Interviews will be analysed using combined deductive framework and inductive thematic analysis.

Intervention Type

Mixed

Primary outcome measure

1. Maternal mortality:
 - 1.1. Maternal deaths
2. Newborn mortality:
 - 2.1. Stillbirths
 - 2.2. Neonatal deaths
3. Maternal morbidity:
 - 3.1. (Pre)-eclampsia
 - 3.2. Uterine rupture
 - 3.3. Antepartum haemorrhage
 - 3.4. Postpartum haemorrhage
 - 3.5. Maternal sepsis
4. Newborn morbidity:
 - 4.1. Birth asphyxia
 - 4.2. Prematurity (born before 37 weeks gestation)
 - 4.3. Low birthweight (weight less than 2500 g)
5. Emergency obstetric referrals
6. Appropriate emergency referrals

Measured by monthly collection of routine health facility records for maternity ward records from the baseline phase to the end of cluster participation.

In addition to the primary and secondary outcomes listed here, the researchers may evaluate general process outcomes as described in the protocol attached.

Secondary outcome measures

1. Proportion of women with the following obstetric complications:
 - 1.1. (Pre)-eclampsia

1.2. Anaemia (Hb less than 7g/dl)

1.3. Syphilis

2. Proportion of women referred out for any cause, or due to pre/eclampsia, anaemia and risk for obstruction of labour

3. Proportion of newborns with the following complications: neonatal sepsis

4. Proportion of neonatal unit (NNU) admission for any cause

5. Proportion of newborn breastfeeding initiated within 60 minutes of birth

6. Proportion Kangaroo Mother Care (KMC) admissions

7. Proportion of newborns resuscitated

8. Proportion of newborns received antibiotics

9. Proportion of women starting antenatal care (ANC) visit in the first trimester (<12 weeks)

10. Average total ANC visits per woman

11. Proportion of women who received required doses of the following (data from registers):

11.1. Tetanus toxoid vaccination (TTV) (≥ 2 doses)

11.2. Sulphadoxine-pyrimethamine (SP) (≥ 3 doses)

Measured by monthly collection of routine health facility maternity and neonatal records and cohort records (6 months prior to reporting month) for ANC data records, from the baseline to the end of cluster participation.

In addition to the primary and secondary outcomes listed here, the researchers may evaluate general process outcomes as described in the protocol attached.

Overall study start date

01/08/2022

Completion date

31/07/2026

Eligibility

Key inclusion criteria

Clusters:

Primary Health Centre facilities offering maternity care will be included as a cluster following the completion of a successful feasibility report with the minimum prerequisites of:

1. A minimum of 700 ANC initial visits to a maximum of 2000 per year
2. Providers of basic emergency obstetric care
3. Public primary facility
4. Completed the site readiness assessment process

Participants: clients (in process evaluation interviews and survey):

1. ≤ 20 weeks of gestation at the initial visit to ANC
2. Able to provide informed consent
3. Willing to participate in the study
4. Aged 18 years and above

Participants: HCWs (in process evaluation interviews and survey):

1. Healthcare workers responsible for the care of women during pregnancy
2. Willing to participate in the study
3. Able to provide informed consent

Participant type(s)

Patient, Health professional

Age group

Adult

Lower age limit

18 Years

Upper age limit

100 Years

Sex

Both

Target number of participants

32 total facilities (clusters) with at least 1600 women per trial arm

Key exclusion criteria

Clusters:

1. PHC with less than 700 initial ANC visits per year
2. PHCs with more than 2000 initial ANC visits per year-outliers to other facilities
3. PHCs not willing to participate in the study
4. High risk of contamination (geographically close to another participating centre or site where similar interventions are already taking place)

Participants: clients (in process evaluation - interviews and survey):

1. >20 weeks of gestation at the initial visit to ANC
2. Unable to provide informed consent
3. Medical and obstetric complications on the initial visit
4. Aged less than 18 years

Participants: HCWs (in process evaluation - interviews and survey):

1. Not willing to participate in process evaluation
2. Unable to provide informed consent

Date of first enrolment

01/08/2025

Date of final enrolment

31/05/2026

Locations**Countries of recruitment**

Malawi

Zambia

Study participating centre

Katchale, Ming'ongo, Chimbalanga, Matapila, Chadza, Ndaula, Malembo, Diamphwe, Maluwa, Chiunjiza, Mbang'ombe2, Mbabzi, Ngoni, Chiwamba, Chiwoza, Nthondo, Nsara, Kang'oma and Khongoni

Lilongwe District Health Centres

Lilongwe

Malawi

207201

Study participating centre

Phirilongwe, Malukula, Namkumba, Chilonga, Kukalanga, Namwera, Mkumba, Malombe, Mtimabii, Lungwena, Chilipa, Nangalamu, Jalasi and Makanjira

Mangochi District Health Centres

Mangochi

Malawi

301400

Study participating centre

Chipapa and Chinyanya Health Centres

Kafue District Health Centres

Kafue

Zambia

None

Study participating centre

Kanakantapa Health Centre

Chongwe District

Chongwe Lusaka

Zambia

None

Study participating centre

Malombe and Chisamba Health Centres

Chisamba District Health Centres

Chisamba

Zambia

None

Study participating centre

Lungobe, Myooye and Kamilambo Health Centres

Mumbwa District Health Centres

Mumbwa
Zambia
None

Sponsor information

Organisation

University of Liverpool

Sponsor details

Institute of Life Course and Medical Sciences
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Sponsor type

Research organisation

Website

<http://liverpool.ac.uk>

ROR

<https://ror.org/04xs57h96>

Funder(s)

Funder type

Government

Funder Name

National Institute for Health and Care Research

Alternative Name(s)

National Institute for Health Research, NIHR Research, NIHRresearch, NIHR - National Institute for Health Research, NIHR (The National Institute for Health and Care Research), NIHR

Funding Body Type

Government organisation

Funding Body Subtype

National government

Location
United Kingdom

Results and Publications

Publication and dissemination plan

Intention to publish date
01/03/2027

Individual participant data (IPD) sharing plan
The data-sharing plans for the current study are unknown and will be made available at a later date. The researchers expect the data sets generated from this study to be available upon request.

IPD sharing plan summary
Available on request

Study outputs

Output type	Details	Date created	Date added	Peer reviewed?	Patient-facing?
Protocol file	version 2.0	12/11/2024	12/03/2025	No	No