

Measuring levels of maternal cell microRNAs to predict adverse pregnancy outcome

Submission date 25/12/2017	Recruitment status No longer recruiting	<input checked="" type="checkbox"/> Prospectively registered
		<input type="checkbox"/> Protocol
Registration date 09/01/2018	Overall study status Completed	<input type="checkbox"/> Statistical analysis plan
		<input type="checkbox"/> Results
Last Edited 01/07/2021	Condition category Pregnancy and Childbirth	<input type="checkbox"/> Individual participant data
		<input type="checkbox"/> Record updated in last year

Plain English summary of protocol

Background and study aims

The aim of this study is to develop a first trimester predictor test for identifying pregnant women at high risks of various pregnancy complications. These complications include high blood pressure in pregnancy (known as pre-eclampsia), miscarriage (pregnancy loss) and spontaneous pre-term birth (premature delivery from 24 to 36 weeks of pregnancy). This predictor test will measure the levels of specific microRNAs from a blood test. microRNAs are small molecules which contain genetic material that is essential for all known forms of life. They control gene expression and regulate many proteins. High levels of miRNAs have been shown to decrease certain gene expression pathways believed to play a role in the development of a healthy pregnancy. This study may help to identify pregnant women who are at high risk of developing pregnancy complications. The outcome of this study may alter the management of future pregnant women deemed to be at risk of developing these adverse outcomes.

Who can participate?

Pregnant women aged 24 to 43

What does the study involve?

Participants provide blood samples at 8-10 weeks of pregnancy. Also, if the participant chooses to return for NIPT (non-invasive prenatal testing) where a blood test is routinely drawn, a very small volume of blood is also drawn for the purpose of this study. The levels of specific miRNAs are measured. Participants are asked to fill out a questionnaire detailing their pregnancy after 9 months. The history of the pregnancy is then compared to the level of miRNAs in the blood.

What are the possible benefits and risks of participating?

There are no direct benefits to those participating. However, the information obtained from this study may help to identify patients who are at high risks of pre-eclampsia, miscarriage or spontaneous preterm delivery. This may result in a change in the management of pregnant women who are identified at risk of adverse pregnancy, and minimise the risk of complications. Drawing blood for normal screening does not carry any risks and participating in this study is not expected to add any further risks.

Where is the study run from?

The Centre for Reproductive and Genetic Health (UK)

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

December 2017 to August 2025

Who is funding the study?

The Centre for Reproductive and Genetic Health (UK)

Who is the main contact?

1. Ms Jara Ben-Nagi (scientific)
2. Miss Rabi Odia (public)

Contact information

Type(s)

Scientific

Contact name

Ms Jara Ben-Nagi

Contact details

230-232 Great Portland Street
London
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Type(s)

Public

Contact name

Miss Rabi Odia

Contact details

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Additional identifiers

Protocol serial number

miRNA

Study information

Scientific Title

Measuring levels of maternal cell microRNAs to predict adverse pregnancy outcome: a longitudinal observational study

Study objectives

Can levels of maternal microRNAs become a first trimester “predictor” test for identifying high risk of an adverse pregnancy outcome (including pre-eclampsia, miscarriage, etc)?

Ethics approval required

Old ethics approval format

Ethics approval(s)

Not provided at time of registration

Study design

Single-centre longitudinal observational study

Primary study design

Observational

Study type(s)

Diagnostic

Health condition(s) or problem(s) studied

Couples that have successfully conceived

Interventions

Peripheral blood is drawn from a pregnant patient at 8-10 weeks, the sample undergoes separation/extraction of PBMCs which is stored at -80°C. miRNA isolation, miRNA quantification, reverse transcription, and real-time PCR will be carried out. Patients who have participated in this study are then asked to fill out a questionnaire detailing their pregnancy after 9 months. The history of pregnancy will then be compared to the level of microRNA in peripheral blood.

1. Sample collection: After obtaining patient consent. Peripheral blood from successfully conceived patients at 8-10 weeks will be collected in Heparin tubes (green top) and incubated at room temperature for 1-2 hours
2. Separation of PBMCs from peripheral blood and storage: Separation/extraction of PBMCs will be performed at room temperature using Ficollhpaque density gradient centrifugation. The PBMCs layer will be “pipetted” into tubes containing 1ml of Trizol and stored at -80oC
3. Molecular techniques: miRNA isolation, miRNA quantification, reverse transcription, and real-time PCR will be performed by igenomix UK
4. Analysis, statistics and reporting: Will be performed by Igenomix UK scientific staff
5. Follow up: Collection of patient birth and adverse outcomes will be conducted by CRGH. Once the agreed endpoint is reached, the sample IDs will be unblinded and the results shared between CRGH and Igenomix UK in preparation for publication

Intervention Type

Other

Primary outcome(s)

Levels of specific miRNAs isolated from maternal peripheral blood mononuclear cells (PBMCs) in 8-10-week pregnant patients, measured by real-time PCR using TaqMan probes after reverse transcription

Key secondary outcome(s)

Pregnancy outcome assessed using a questionnaire after 9 months

Completion date

01/08/2025

Eligibility

Key inclusion criteria

Successfully pregnant healthy females aged between 24 and 43

Participant type(s)

Healthy volunteer

Healthy volunteers allowed

No

Age group

Adult

Lower age limit

24 years

Upper age limit

43 years

Sex

Female

Key exclusion criteria

Under 24 and over 43 years old females who are pregnant

Date of first enrolment

30/01/2018

Date of final enrolment

30/09/2024

Locations

Countries of recruitment

United Kingdom

England

Study participating centre

The Centre for Reproductive and Genetic Health

230-232 Great Portland Street

London

United Kingdom
W1W 5QS

Sponsor information

Organisation

Centre for Reproductive and Genetic Health

Funder(s)

Funder type

Hospital/treatment centre

Funder Name

Centre for Reproductive and Genetic Health

Results and Publications

Individual participant data (IPD) sharing plan

The datasets generated during and/or analysed during the current study are/will be available upon request from the Centre for Reproductive and Genetics Health (jarabennagi@crgh.co.uk). Data will be available upon completion of the study for a minimum of 5 years. Data will be accessed via an internal shared drive. Access to the shared drive has a designated username and password. The access will only be to the designated research members via an NHS secure web if receiving party also has access to this and password protected. This will be shared with iGenomix UK and only be available upon request. Consent will be obtained from the patients. The data will only contain participants' internal reg number and is password protected.

IPD sharing plan summary

Available on request