

Assessing the effects of maternal seizures during pregnancy on the brain development of children

| | | |
|----------------------------------------|------------------------------------------------------|--------------------------------------------------------------------------------------------------------------|
| Submission date 14/12/2018 | Recruitment status No longer recruiting | <input checked="" type="checkbox"/> Prospectively registered <input type="checkbox"/> Protocol |
| Registration date 09/01/2019 | Overall study status Completed | <input type="checkbox"/> Statistical analysis plan <input type="checkbox"/> Results |
| Last Edited 09/07/2020 | Condition category Nervous System Diseases | <input type="checkbox"/> Individual participant data <input type="checkbox"/> Record updated in last year |

Plain English summary of protocol

Background and study aims

The treatment of epilepsy during pregnancy is a balance between treating the seizures and reducing, as far as possible, the risks presented to the developing child by medication. This study will investigate whether seizures which the mother has during pregnancy alters the brain development of the developing child.

Who can participate?

Women with epilepsy who are either pregnant (and not yet 21 weeks) or women who are trying for a baby currently. Women must be taking the medications lamotrigine (lamictal) or levetiracetam (keppra) alone or alongside other antiepileptic medications (but not if they are taking sodium valproate (epilim) or carbamazepine (tegtretol)). The study is looking for volunteers both who are having seizures and those who are not having seizures.

What does the study involve?

All participants will be asked to sign a consent form indicating that they understand the study and are happy for themselves and their baby to participate. They will then be asked to provide some information about their health, education, occupation and to record on a daily basis whether or not they have had seizure. Women can do this either in a paper seizure diary or using an app. When the women's pregnancy has passed 32 weeks they will be asked about their general health and the medications they are taking again. When the baby is born researchers at the local hospital will record information about the babies birth from medical records. Shortly after the child's first birthday a home visit will be completed where a play-based assessment will be completed with the child. This involves playing a number of games with the child, which in turn provides information on their development in areas such as reasoning, language and motor abilities. Their mothers will also be asked to complete a questionnaire about their early social development and about the mothers own mood. Mother's will also be asked to complete two brief tasks (e.g. one puzzle and one language task) to inform on their reasoning abilities.

What are the possible benefits and risks of participating?

There may be no direct benefits personally to taking part. However, following the assessment of

the child feedback would be provided to parents and a letter be sent the child's GP and kept their medical record. If the person who completed the assessment had concerns about specific areas of the child's development they would discuss it with the parents and then send a letter to the child's GP. In the longer term, the findings of this study will be of benefit to both women with epilepsy and their doctors by providing information about how seizures in pregnancy may or may not impact on how the child develops in their first year of life. This may be directly relevant to participants in the study if they were planning another pregnancy in the future.

If the participant does not already record her seizures it might seem a big task to record them everyday, however only a few details are needed and on most days the participant may simply be ticking 'no' if they did not have any seizures. A further possible disadvantage of this study is that the developmental assessment may reveal that the child is experiencing some difficulties in one or more areas of their development. In this situation, with parental permission, we would share this information with the child's GP to make sure that they can arrange the necessary support for the child.

Where is the study run from?

The study is being led by researchers at the University of Manchester and Manchester University Hospitals NHS Foundation Trust but involves doctors and nurses from 21 hospitals around the UK.

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

Set up for the study started in May 2018 and it will open to recruitment on the 14th January 2018. Women will be able to opt into the study until May 2020 and after this time no new participants will be able to opt into the study. The follow-ups after the child's first birthday will run until August 2021. The study will close on 30th October 2021 with the results being available shortly after that.

Who is funding the study?

Epilepsy Research UK

Who is the main contact?

Myfanwy Rawson

Myfanwy.rawson@mft.nhs.uk

Contact information

Type(s)

Public

Contact name

Ms Myfanwy Rawson

Contact details

Genomic Medicine Research Office

St Mary's Hospital

Oxford Road

Manchester

United Kingdom

M13 9WL
0161 701 2788
myfanwy.rawson@mft.nhs.uk

Type(s)
Scientific

Contact name
Dr Rebecca Bromley

ORCID ID
<http://orcid.org/0000-0003-4008-0917>

Contact details
School of Biological Sciences
University of Manchester
Manchester
United Kingdom
M13 9WL
0161 701 4514
rebecca.bromley@manchester.ac.uk

Additional identifiers

EudraCT/CTIS number

IRAS number
232507

ClinicalTrials.gov number

Secondary identifying numbers
IRAS 232507

Study information

Scientific Title
Neurodevelopment After Prenatal Exposure to Seizures study: a cohort study

Acronym
NAPES

Study objectives
Children exposed to frequent seizures in the womb will have poorer development at 12 months of age.

Ethics approval required
Old ethics approval format

Ethics approval(s)

Study design

Multi-centre prospective observational cohort study.

Primary study design

Observational

Secondary study design

Cohort study

Study setting(s)

Hospital

Study type(s)

Other

Participant information sheet

Not available in web format. Please request via email rebecca.bromley@manchester.ac.uk

Health condition(s) or problem(s) studied

Epilepsy

Interventions

Participants are recruited from their local participating hospital. Following recruitment information will be taken from participants regarding their background, such as their education, health and occupation. From this point they record their seizures on a daily basis using either a paper seizure diary or an app. Each month their local research team will contact them to collect that month's seizure information. In the third trimester (after 32 weeks) the participant will be contacted to provide information about their current health and medications which they are taking. Following the birth of the child information is recorded about the delivery method and the health of the child at birth. After the child's first birthday, an appointment will be arranged to see the child and parent at home where a play-based assessment will be completed. Whilst this is fun for the child it provides the assessor with important information about where the child is up to in terms of their cognitive (reasoning), language and motor development. Following the assessment of development a questionnaire will be posted to participants which will ask them about their experience of taking part in the study.

Intervention Type

Other

Primary outcome measure

1. Cognitive development is measured using Bayley Scales of Infant and Toddler Development at 12- 15 months of age.
2. Language development is measured using Bayley Scales of Infant and Toddler Development at 12- 15 months of age.
3. Motor development is measured using Bayley Scales of Infant and Toddler Development at 12- 15 months of age.

Secondary outcome measures

1. Social and behavioural development is measured using Bayley Scales of Infant and Toddler Development at 12- 15 months of age.
2. Maternal experiences of participating in the study is measured using questionnaires when the child is 12- 15 months of age.

Overall study start date

18/05/2018

Completion date

30/04/2022

Eligibility

Key inclusion criteria

1. They have a diagnosis of epilepsy and are either:
 - 1.1. Trying to conceive
 - 1.2. They are pregnant and before 21 weeks gestation
 - 1.2.1. They have an expected date of delivery prior to the recruitment closure date (30th November 2019).
2. They are taking either:
 - 2.1. Monotherapy lamotrigine or levetiracetam
 - 2.2. Polytherapy combinations including lamotrigine and levetiracetam and other antiepileptic drugs; with the exception of valproate and carbamazepine.
3. They are receiving their care through one of the participating centres
4. They are willing to complete prospectively recorded seizure diaries

Participant type(s)

Patient

Age group

Adult

Sex

Female

Target number of participants

180

Key exclusion criteria

1. Significant learning disability, requiring support to live independently
2. Experience or history of non-epileptic attacks (also known as NEAD or psychogenic seizures)
3. Taking non-antiepileptic medications which are known to be teratogenic (e.g. warfarin, mycophenolate)
4. English is not the first language in the home

Date of first enrolment

14/01/2019

Date of final enrolment

30/03/2021

Locations

Countries of recruitment

England

Northern Ireland

United Kingdom

Study participating centre

Manchester University NHS Foundation Trust

Oxford Road

Manchester

United Kingdom

M13 9WL

Study participating centre

Belfast Health and Social Care Trust

Royal Hospitals Grosvenor Road

Belfast

United Kingdom

BT12 6BA

Study participating centre

Lancashire Teaching Hospitals NHS Foundation Trust

Royal Preston Hospital

Sharoe Green Lane

Preston

United Kingdom

PR2 9HT

Study participating centre

Liverpool Women's Hospital NHS Foundation Trust

Crown Street

Liverpool

United Kingdom

L8 7SS

Study participating centre

Salford Royal NHS Foundation Trust

Stott Lane
Salford
United Kingdom
M6 8HD

Study participating centre

City Hospitals Sunderland NHS Foundation Trust

Sunderland Royal Hospital
Kayla Road
Sunderland
United Kingdom
SR4 7TR

Study participating centre

Newcastle Upon Tyne Hospitals NHS Foundation Trust

Royal Victoria Hospital
Queen Victoria Road
Newcastle Upon Tyne
United Kingdom
NE1 4LP

Study participating centre

South Tees Hospitals NHS Foundation Trust

Marlon Road
Middlesbrough
United Kingdom
TS4 3BW

Study participating centre

Walton Centre for Neurology and Neurosurgery NHS Foundation Trust

Lower Lane
Liverpool
United Kingdom
L9 7LJ

Study participating centre

University Hospitals of Morecambe Bay NHS Foundation Trust

Lancaster Royal Infirmary
Ashton Road
Lancaster

United Kingdom
LA1 4RP

Study participating centre

Warrington and Halton Hospitals NHS Foundation Trust

Lovely Lane
Warrington
United Kingdom
WA5 1QG

Study participating centre

Countess of Chester Hospital NHS Foundation Trust

The Countess Of Chester Health Park
Liverpool Road
Chester
United Kingdom
CH2 1UL

Study participating centre

East Lancashire Hospitals NHS Foundation Trust

Royal Blackburn Hospital
Haslingden Road
Blackburn
United Kingdom
BB2 3HH

Study participating centre

Mid Cheshire Hospitals NHS Foundation Trust

Leighton Hospital
Crewe
United Kingdom
CW1 4QJ

Study participating centre

York Teaching Hospitals NHS Foundation Trust

Wigginton Road
York
United Kingdom
YO31 8HE

Study participating centre

Leeds Teaching Hospitals NHS Foundation Trust

St James Hospital

Beckett Street

Leeds

United Kingdom

LS9 7TF

Study participating centre

University Hospitals Birmingham NHS Foundation Trust

Queen Elizabeth Hospital Birmingham

Mindelsohn Way

Birmingham

United Kingdom

B15 2GW

Study participating centre

County Durham and Darlington NHS Foundation Trust

North Road

Durham

United Kingdom

DH1 5TW

Study participating centre

Blackpool Teaching Hospitals NHS Foundation Trust

Blackpool Victoria Hospital

Whinney Heys Road

Blackpool

United Kingdom

FY3 8NR

Study participating centre

Stockport NHS Foundation Trust

Stepping Hill Hospital

Stockport

United Kingdom

SK2 7JE

Study participating centre

The Mid Yorks Hospital NHS Foundation Trust
Pinderfields Hospital
Wakefield
United Kingdom
WF1 4DG

Sponsor information

Organisation

University of Manchester

Sponsor details

Oxford Road
Manchester
England
United Kingdom
M13 9PL
0161306 0677
FBMHethics@manchester.ac.uk

Sponsor type

University/education

ROR

<https://ror.org/027m9bs27>

Funder(s)

Funder type

Charity

Funder Name

Epilepsy Research UK

Alternative Name(s)

ERUK

Funding Body Type

Private sector organisation

Funding Body Subtype

Other non-profit organizations

Location

Results and Publications

Publication and dissemination plan

The results will be shared with participants and will be written up for formal scientific publication. We will also undertake a number of local, national and international talks to share the research findings.

Intention to publish date

30/10/2022

Individual participant data (IPD) sharing plan

The datasets generated during and/or analysed during the current study are/will be available upon request from Dr Rebecca Bromley, rebecca.bromley@manchester.ac.uk, in electronic format, within a year of the study end for utilisation in studies with similar aims and where the participant has consented.

IPD sharing plan summary

Available on request

Study outputs

| Output type | Details | Date created | Date added | Peer reviewed? | Patient-facing? |
|--------------------------------------|---------|--------------|------------|----------------|-----------------|
| HRA research summary | | | 28/06/2023 | No | No |