

Thyroid AntiBodies and LEvoThyroxine study (TABLET)

Submission date 31/10/2011	Recruitment status No longer recruiting	☒ Prospectively registered ☐ Protocol
Registration date 31/10/2011	Overall study status Completed	☐ Statistical analysis plan ☒ Results
Last Edited 28/05/2020	Condition category Pregnancy and Childbirth	☐ Individual participant data

Plain English summary of protocol
Not provided at time of registration

Contact information

Type(s)
Scientific

Contact name
Prof Arri Coomarasamy

Contact details
Professor of Gynaecology
Director, Tommy's National Centre for Miscarriage Research
Institute of Metabolism and Systems Research
University of Birmingham, UK
Consultant Gynaecologist and Sub-specialist in Reproductive Medicine
Birmingham Women's Hospital, UK
Birmingham
United Kingdom
B15 2TG
+44 (0)121 627 2775
s.j.magson@bham.ac.uk

Additional identifiers

Clinical Trials Information System (CTIS)
2011-000719-19

Protocol serial number
11051

Study information

Scientific Title

A randomised controlled trial of the efficacy and mechanism of levothyroxine treatment on pregnancy and neonatal outcomes in women with thyroid antibodies

Acronym

TABLET

Study objectives

Current study hypothesis as of 29/04/2013:

Levothyroxine will assist women aged 16-40 with a history of one or more miscarriage or who are being treated for infertility, who have been identified as having thyroid antibodies (TPO antibody positive) trying to conceive within 1 year of miscarriage.

Previous study hypothesis until 29/04/2013:

Levothyroxine will assist women aged 16-40 with a history of one or more miscarriage, who have been identified as having thyroid antibodies (TPO antibody positive) trying to conceive within 1 year of miscarriage.

Ethics approval required

Old ethics approval format

Ethics approval(s)

South West C REC, 29/03/2011, ref: 11/SW/0036

Primary study design

Interventional

Study design

Randomised interventional trial

Study type(s)

Treatment

Health condition(s) or problem(s) studied

Reproductive health, childbirth

Interventions

Current interventions as of 29/04/2013:

Women aged 16-40 with a history of one or more miscarriage or who are being treated for infertility, who have been identified as having thyroid antibodies (TPO antibody positive), and will be trying to conceive within 1 year.

Previous interventions until 29/04/2013:

Women aged 16-40 with a history of one or more miscarriage, who have been identified as having thyroid antibodies (TPO antibody positive), and will be trying to conceive within 1 year of miscarriage.

Levothyroxine, 50mcg Levothyroxine, overencapsulated to be taken orally daily versus placebo, to match investigational medicinal products (IMP). Followed Up at 21 month(s)

Intervention Type

Drug

Phase

Phase III

Drug/device/biological/vaccine name(s)

Levothyroxine

Primary outcome(s)

Current primary outcome measures as of 14/09/2017:

Live birth at or beyond 34 weeks (greater than or equal to 34). The denominator of this proportion will be all women randomised, and the numerator will be those women who proceed to have a live birth at or beyond 34 weeks.

Previous primary outcome measures:

Live birth measured at beyond 34 weeks gestational age

Key secondary outcome(s)

Current secondary outcome measures as of 14/09/2017:

1. Clinical pregnancy assessed using urine test at 7 weeks
2. Ongoing pregnancy assessed using scan at 12 weeks
3. Miscarriage <24 weeks, clinical assessment
4. Stillbirth (inter-uterine death ≥ 24 weeks), clinical assessment
5. Ectopic pregnancy, clinical assessment
6. Termination (and reasons), clinical assessment
7. Live birth <34 weeks, clinical assessment
8. Time from conception to pregnancy end (any reason)
9. Mode of initiation of labour (spontaneous/induced)
10. Mode of delivery (vaginal/operative vaginal/caesarean)
11. Gestation at delivery, weeks (scan)
12. Time from conception to live birth
13. Gestation at delivery <28 weeks/<34 weeks/<37 weeks
14. Birth weight, grams
15. Birth weight adjusted for gestational age and sex, centiles
16. Birth weight adjusted for gestational age, sex, parity, maternal BMI and ethnicity, centiles
17. Small for gestational age and sex (birth weight proportion <10th centile)
18. Small for gestational age, sex, parity, maternal BMI and ethnicity (birth weight proportion <10th centile)
19. Large for gestational age and sex (birth weight proportion ≥ 90 th centile)
20. Large for gestational age, sex, parity, maternal BMI and ethnicity (birth weight proportion ≥ 90 th centile)
21. APGAR score, clinical assessment at 1 minute/5 minutes
22. Serum TSH concentration ($\mu\text{U/L}$; log transformed), assessed by blood test at each assessment time (3 months, 6 months, 9 months, 6-8 weeks pregnancy, 16-18 weeks pregnancy and 28 weeks pregnancy)
23. Serum free T4 level (pmol/L), assessed by blood test at each assessment time (3 months, 6

months, 9 months, 6-8 weeks pregnancy, 16-18 weeks pregnancy and 28 weeks pregnancy)
24. Maternal antenatal complications (hyperemesis gravidarum/gestational diabetes/pre-eclampsia or eclampsia/obstetric cholestasis/pre-term pre labour rupture of membranes (PPROM)/intrauterine growth restriction (IUGR)/others), clinical assessment
25. Intrapartum complications (shoulder dystocia/others), clinical assessment
26. Maternal postnatal complications (admission to HDU or ITU/Abnormal thyroid test within four weeks/referred to psychiatrist or started on antidepressants/others), clinical assessment
27. Neonatal complications (early neonatal death defined as death within 7 days after delivery /late neonatal death defined as death beyond 7 days and before 28 days post-delivery/admission to NNU/SCBU/active resuscitation within first 28 days/surfactant use/mechanical ventilation /intermittent positive pressure ventilation/continuous positive airway pressure/oxygen use /congenital abnormalities/hypoxic ischaemic encephalopathy/retinopathy of prematurity /respiratory distress syndrome/pneumothorax/intraventricular hemorrhage (grade 3 or 4) /necrotizing enterocolitis/early infection/others), clinical assessment
28. Reported symptoms that participant is concerned about at each assessment time (3 months, 6 months, 9 months, 6-8 weeks pregnancy, 16-18 weeks pregnancy and 28 weeks pregnancy)
29. Serious Adverse Events, clinical assessment at 3 months, 6 months, 9 months, 6-8 weeks pregnancy, 16-18 weeks pregnancy and 28 weeks pregnancy

Previous secondary outcome measures (added 29/04/2013):

1. Gestation at delivery
2. Conception (urinary pregnancy) rate
3. Clinical pregnancy at 6-8 weeks
4. On-going pregnancy at 11-13 weeks
5. Miscarriage (delivery before 24 weeks of gestation)
6. Survival at 28 days of neonatal life
7. Thyroid function tests

Completion date

15/05/2017

Eligibility

Key inclusion criteria

Current inclusion criteria as of 29/04/2013:

1. Women trying to conceive
2. History of one or more miscarriage(s) or who are being treated for infertility
3. Age 16-40 years at randomisation
4. Biochemically euthyroid (Free T4 and TSH within specified reference ranges)
5. Thyroid Peroxidase Antibody (TPO) positive
6. Willing and able to give written informed consent

Previous inclusion criteria until 29/04/2013:

1. Women trying to conceive
2. History of one or more miscarriage(s)
3. Age 16-40 years at randomisation
4. Biochemically euthyroid (Free T4 and TSH within specified reference ranges)
5. Thyroid Peroxidase Antibody (TPO) positive
6. Willing and able to give written informed consent

Participant type(s)

Patient

Healthy volunteers allowed

No

Age group

Adult

Sex

Female

Total final enrolment

952

Key exclusion criteria

Current exclusion criteria as of 29/04/2013:

1. Current or past treatment for thyroid disease
2. Contraindication to levothyroxine treatment
3. Thyrotoxicosis
4. Hypersensitivity to any components tablets
5. Women taking amiodarone or lithium therapy
6. Participants in any other blinded, placebo controlled trials on Investigational Medicinal Products in pregnancy
7. Previous or current diagnosis of cardiac disease

Previous exclusion criteria until 29/04/2013:

1. Current or past treatment for thyroid disease
2. Contraindication to levothyroxine treatment
3. Thyrotoxicosis
4. Hypersensitivity to any components tablets
5. Women taking amiodarone or lithium therapy
6. Participants in any other blinded, placebo controlled trials on Investigational Medicinal Products in pregnancy
7. Planning to conceive using ovulation stimulation therapy
8. Previous or current diagnosis of cardiac disease

Date of first enrolment

01/11/2011

Date of final enrolment

22/01/2016

Locations

Countries of recruitment

United Kingdom

England

Study participating centre
Birmingham Womens Hospital
Birmingham
United Kingdom
B15 2TG

Study participating centre
For full list of centres please see: <http://www.birmingham.ac.uk/research/activity/mds/trials/bctu/trials/womens/tablet/investigators/recruitment.aspx>
United Kingdom
-

Sponsor information

Organisation
University of Birmingham (UK)

ROR
<https://ror.org/03angcq70>

Funder(s)

Funder type
Government

Funder Name
Efficacy and Mechanism Evaluation Programme

Alternative Name(s)
NIHR Efficacy and Mechanism Evaluation Programme, Efficacy and Mechanism Evaluation (EME), EME

Funding Body Type
Government organisation

Funding Body Subtype
National government

Location
United Kingdom

Results and Publications

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

IPD sharing plan summary

Data sharing statement to be made available at a later date

Study outputs

Output type	Details	Date created	Date added	Peer reviewed?	Patient-facing?
Results article	results	04/04/2019		Yes	No
Basic results			28/05/2020	No	No
HRA research summary			28/06/2023	No	No
Participant information sheet	Participant information sheet	11/11/2025	11/11/2025	No	Yes
Study website	Study website	11/11/2025	11/11/2025	No	Yes