

Exposure to nicotine and tobacco products during pregnancy

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| Submission date 25/11/2019 | Recruitment status No longer recruiting | <input type="checkbox"/> Prospectively registered <input type="checkbox"/> Protocol |
| Registration date 27/11/2019 | Overall study status Completed | <input type="checkbox"/> Statistical analysis plan <input type="checkbox"/> Results |
| Last Edited 17/01/2025 | 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

Health professionals and pregnant women are cautious about recommending or using e-cigarettes (ECs) in pregnancy, due to lack of detailed safety information. The main aim of the study is to measure toxins and cancer causing substances in the urine of women who used to smoke and now just use ECs compared with women who just smoke.

Who can participate?

Pregnant women who are smokers, users of ECs, nicotine replacement therapy, smoke and use ECs, or have never smoked.

What does the study involve?

Pregnant women will be invited to take part when they attend a routine antenatal appointment. If they agree to participate, they will be asked by a researcher to provide a urine sample (or a routinely collected sample will be used if available) and to complete a questionnaire (about themselves and use of nicotine/tobacco products, alcohol, cannabis). They will also be asked to blow into a device that measures exposure to tobacco smoke. They will then be asked to meet with the researcher again at a further routine antenatal visit, in order to repeat the assessments.

What are the possible benefits and risks of participating?

None

Where is the study run from?

1. St George's University Hospitals NHS Foundation Trust, UK
2. Lancashire teaching Hospitals NHS Foundation Trust, UK
3. Bradford Teaching Hospital NHS Foundation Trust, UK
4. The Newcastle Upon Tyne Hospitals NHS Foundation Trust, UK
5. Gateshead Health NHS Foundation Trust, UK

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

November 2019 to October 2021

Who is funding the study?
Cancer Research UK

Who is the main contact?
Prof. Michael Ussher
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Contact information

Type(s)
Scientific

Contact name
Prof Michael Ussher

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Additional identifiers

Clinical Trials Information System (CTIS)
Nil known

Integrated Research Application System (IRAS)
269631

ClinicalTrials.gov (NCT)
Nil known

Protocol serial number
V4.0; IRAS project ID: 269631

Study information

Scientific Title
Maternal exposure to carcinogens, toxicants and nicotine during pregnancy among e-cigarette users, smokers, nicotine replacement therapy users and among those who have never used nicotine or tobacco products

Study objectives

During pregnancy, we hypothesise that electronic cigarette (EC) users will have significantly lower levels of urinary acrolein than smokers. As secondary objectives we will explore levels of a range of biomarkers in exclusive EC users compared with exclusive smokers, dual users of cigarettes and ECs, exclusive nicotine replacement therapy (NRT) users; dual users of cigarettes and NRT, and 'never users of tobacco or nicotine products'. Generally, the hypothesis is that less tobacco smoke exposure will equate to less toxin and carcinogen exposure.

Ethics approval required

Old ethics approval format

Ethics approval(s)

Approved 29/10/2019, London - Brighton & Sussex Research Ethics Committee (Health Research Authority, Skipton House, 80 London Road, London, SE1 6LH; +44(0)207 104 8052; NRESCommittee.SECOast-BrightonandSussex@nhs.net), ref: 19/LO/1599

Study design

Observational cross-sectional multicentre study

Primary study design

Observational

Study type(s)

Other

Health condition(s) or problem(s) studied

Use of nicotine and tobacco products during pregnancy

Interventions

Pregnant women will be invited to take part when they attend a routine antenatal appointment. If they agree to participate, they will be asked by a researcher to provide a urine sample (or a routinely collected sample can be used if available) and to complete a questionnaire (about themselves and use of nicotine/tobacco products, alcohol, cannabis). They will also be asked to blow into a device that measures exposure to tobacco smoke. They will then be asked to meet with the researcher again at a further routine antenatal visit, in order to repeat the assessments. Urine samples will be stored in a freezer and at the end of the study will be sent to an external laboratory for testing. The results will provide valuable information for women who are pregnant, for healthcare professionals who care for those women and for regulators, about the potential safety of using ECs during pregnancy.

Intervention Type

Other

Primary outcome(s)

Current primary outcome measure as of 13/12/2021:

Maternal urinary levels of the volatile organic compound Acrolein (metabolite HPMa) measured using urine test at the time of participation. Acrolein has two metabolites (3HPMA & 2CoEMA); the single primary outcome will be 3HPMA because 3HPMA has a higher ratio in smokers vs non-smokers compared with 2CoEMA. The metabolite 2CoEMA will be analysed as a secondary outcome.

Previous primary outcome measure:

Maternal urinary levels of the volatile organic compound Acrolein (metabolites HPMa and CEMA) measured using urine test at the time of participation

Key secondary outcome(s)

Current secondary outcome measures as of 13/12/2021:

All tested in maternal urine at time of participation:

1. Tobacco exposure will be assessed with a test for minor tobacco alkaloids (anabasine, anatabine), expired carbon monoxide levels
2. Levels of nicotine, cotinine and their metabolites, to derive total nicotine equivalents
3. Nicotine metabolite ratio (NMR)
4. Two tobacco-specific nitrosamines (TSNAs) (i.e., 4-(methylnitrosamino)-1-(3-pyridyl)-1-butanone (NNAL) and N-nitrosonornicotine (NNN)), two heavy metals (i.e., cadmium (Cd) and lead (Pb)) and one polycyclic aromatic hydrocarbon (PAH) (i.e., 2-naphthol)
5. 13 volatile organic compounds (VOCs) parent (metabolite): acrolein (2CoEMA), 1,3-butadiene (MHB3), acrylamide (AAMA), acrylonitrile (CYMA), Benzene (PMA), crotonaldehyde (HPMM), cyanide (ATCA), isoprene (IPM3), ethylbenzene (PHGA), propylene oxide (HPM2), styrene (MADA), toluene/benzyl alcohol (BMA), and m,p-xylene (34MH)
6. Human chorionic gonadotropin (hCG)
7. Pregnancy-associated plasma protein A (PAPPA-A)
8. Use of alcohol and cannabinoids

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6. Human chorionic gonadotropin (hCG)
7. Pregnancy-associated plasma protein A (PAPPA-A)
8. Use of alcohol and cannabinoids

Completion date

31/10/2021

Eligibility

Key inclusion criteria

1. Pregnant women
2. Aged at least 16 years of age
3. Able to speak and read English
4. Do not report having had an active infection (chest infection, cold or flu, sore throat, or fever) within 24 hours of either of the two assessments
5. Willing and able to give informed consent for participation in the study

6. Meet criteria for one of six groups related to level and type of use of nicotine and tobacco products:

6.1. Group A. 'Exclusive smokers': Smoked more than 5 cigarettes every day for more than the last six months and have not vaped or used NRT in the last month

6.2. Group B. 'ECs only': Not smoked cigarettes for at least a month and have vaped every day for at least the last two weeks. Also, have not used NRT in the last month

6.3. Group C. 'NRT only': Not smoked cigarettes for at least a month and have used NRT every day for at least the last two weeks. Also, have not vaped in the last month

6.4. Group D. Dual users of cigarettes and ECs: Smoked cigarettes on most days of the week for more than 6 months. Also, have been using e-cigarettes on most days of the week for at least the last two weeks

6.5. Group E. Dual users of cigarettes and NRT: Smoked cigarettes on most days of the week for more than 6 months. Also, have been using NRT on most days for at least the last two weeks

6.6. Group F. 'never smokers': Have never smoked or used any nicotine or tobacco products

Participant type(s)

Patient

Healthy volunteers allowed

No

Age group

Mixed

Lower age limit

16 years

Sex

Female

Total final enrolment

149

Key exclusion criteria

Current participant exclusion criteria as of 20/10/2021:

1. Have used tobacco products other than cigarettes in the last month (e.g., cigars, heat-not-burn)

2. Have used both NRT and ECs in the last month

3. Have used nicotine free e-cigarettes in the last month

4. Report having had an active infection (chest infection, cold or flu, sore throat, or fever) within 24 hours of either of the two assessments

5. Reported as not currently smoking with an expired carbon monoxide reading of >3 parts per million (where an expired carbon monoxide reading has been obtained)

Previous participant exclusion criteria:

1. Have used tobacco products other than cigarettes in the last month (e.g., cigars, heat-not-burn)

2. Have used both NRT and ECs in the last month

3. Have used nicotine free e-cigarettes in the last month

4. Report having had an active infection (chest infection, cold or flu, sore throat, or fever) within 24 hours of either of the two assessments

Date of first enrolment

14/11/2019

Date of final enrolment

30/04/2020

Locations

Countries of recruitment

United Kingdom

England

Study participating centre

St George's University Hospitals NHS Foundation Trust

Blackshaw Road

London

United Kingdom

SW17 0QT

Study participating centre

Lancashire teaching Hospitals NHS Foundation Trust

Preston Royal Hospital

Sharoe Green Lane

Preston

United Kingdom

PR2 9HT

Study participating centre

Bradford Teaching Hospital NHS Foundation Trust

Bradford Royal Infirmary

Smith Lane

Bradford

United Kingdom

BD9 6RJ

Study participating centre

The Newcastle Upon Tyne Hospitals NHS Foundation Trust

The Royal Victoria Infirmary

Queen Victoria Road

Newcastle upon Tyne
United Kingdom
NE1 4LP

Study participating centre
Gateshead Health NHS Foundation Trust
Queen Elizabeth Hospital
Queen Elizabeth Avenue
Sheriff Hill
Gateshead
United Kingdom
NE9 6SX

Sponsor information

Organisation
St George's University Hospitals NHS Foundation Trust

ROR
<https://ror.org/039zedc16>

Funder(s)

Funder type
Charity

Funder Name
Cancer Research UK

Alternative Name(s)
CR_UK, Cancer Research UK - London, Cancer Research UK (CRUK), CRUK

Funding Body Type
Private sector organisation

Funding Body Subtype
Other non-profit organizations

Location
United Kingdom

Results and Publications

Individual participant data (IPD) sharing plan

The datasets generated during and/or analysed during the current study are not expected to be made available as consent has not be obtained for this purpose.

IPD sharing plan summary

Not expected to be made available

Study outputs

| Output type | Details | Date created | Date added | Peer reviewed? | Patient-facing? |
|---|-------------------------------|--------------|------------|----------------|-----------------|
| HRA research summary | Participant information sheet | | 28/06/2023 | No | No |
| Participant information sheet | | 11/11/2025 | 11/11/2025 | No | Yes |
| Plain English results | | | 17/01/2025 | No | Yes |