ISRCTN10474633 https://doi.org/10.1186/ISRCTN10474633

CARRIAGE study: investigating Staphylococcus aureus nasal carriage

Submission date
11/07/2016

Recruitment status
No longer recruiting

Registration date
20/07/2016

Overall study status
Ongoing

Last Edited
27/02/2023

Condition category
Infections and Infestations

[X] Prospectively registered
[_] Protocol
[_] Statistical analysis plan
[_] Results
[_] Individual participant data
[_] Record updated in last year

Plain English summary of protocol

Background and study aims
Staphylococcus aureus (S. aureus) is a common bacterium with about one person in every three carrying it in their nose, or elsewhere on the skin. Some people carry it all the time, some only now and then, and others never carry it. Currently, we can't explain why. For most people, carriage of S. aureus is harmless. However, a small number of people are susceptible to S. aureus infections due to a weakened immune system and, in some cases, these infections may be serious and resistant to antibiotic treatment. Understanding why only some people are carriers while others aren't will help design new ways to prevent and treat such infections. The aim of this study is to investigate why some people carry S. aureus in their nose while others never do. The study will provide a better understanding of the biological, genetic and environmental factors that determine S. aureus carriage. One way to see if a person carries S. aureus is to analyse swabs taken from the nose.

Who can participate?
Healthy volunteers who participated in the, INTERVAL,COMPARE, and STRIDES BioResource (part of the NIHR BioResource) studies.

What does the study involve?
In phase 1, 250 participants are randomly allocated to one of two swabbing frequencies (two versus three swabs). The acceptability of each swabbing frequency is assessed based on participant response rates and participant feedback from an online questionnaire asking about experiences of taking part in the study.
In phase 2 a further 2000 participants are invited to collect nasal swabs. The number of swabs collected is informed by the results of phase 1 (i.e., either two or three). This phase assesses the feasibility of collecting, processing and analysing swabs from large numbers of participants.
In phase 3, participants from the INTERVAL, COMPARE and STRIDES BioResource studies are invited to collect three nasal swabs so that we receive a full sample set from 20,000 participants in total. This is the phase which follows on from phase 1 and is an extension of phase 2 which tested the feasibility of the study design.

What are the possible benefits and risks of participating?
We will not provide any new information or advice to participants based on the samples and

data collected during the study; at present, this information has no clinical implications for the participant. As such, involvement in the study will have no direct benefit to participants. However, it is an opportunity for volunteers to support research, which addresses an important public health issue. There are no known risks associated with nasal swabbing. All participants will receive clear written instructions and a link to an online video demonstrating swabbing technique. Nasal swabbing may pose minor risks in individuals with nasal problems or nasal damage and, as such, potential participants who report having such conditions will be excluded from the study.

Where is the study run from?
The study is coordinated by the University of Cambridge and is conducted in the community across England.

When is the study starting and how long is it expected to run for?
September 2016 to September 2023

Who is funding the study?
1. Wellcome Trust
2. UK Research and Innovation (UKRI)
3. Health Data Research (HDR) UK
4. The Isaac Newton Trust

Who is the main contact?
Dr Ewan Harrison, eh439@medschl.cam.ac.uk

Study website
http://www.carriagestudy.org.uk

Contact information

Type(s)
Public

Contact name
Dr Ewan Harrison

ORCID ID
http://orcid.org/0000-0003-2720-0507

Contact details
Cardiovascular Epidemiology Unit
Department of Public Health and Primary Care
University of Cambridge
Strangeways Research Laboratory
Worts Causeway
Cambridge
United Kingdom
CB1 8RN
+44 (0)1223 336839
eh439@cam.ac.uk

Additional identifiers

EudraCT/CTIS number
Nil known

IRAS number
202688

ClinicalTrials.gov number
Nil known

Secondary identifying numbers
Protocol version 4 - 20/06/2018, IRAS 202688

Study information

Scientific Title
Understanding the biological basis of persistent carriage of Staphylococcus aureus in humans

Acronym
CARRIAGE

Study objectives
The principal research question is to assess the feasibility and acceptability of protocols for the collection and processing of nasal self-swabs in healthy volunteers to inform a larger proposed study to investigate the biological basis of persistent Staphylococcus aureus (S. aureus) carriage.

The secondary research objectives are to assess:
1. The relative participant-acceptability of two vs three weekly nasal swabs
2. The prevalence of persistent S. aureus carriage rates in healthy volunteers
3. Lifestyle and biomarker correlates of persistent nasal carriage and validate associations with previously suggested candidate variants, and to identify new genetic associations (added 19/06/2019)

Ethics approval required
Old ethics approval format

Ethics approval(s)
North West - Lancaster Research Ethics Committee, 27/06/2016, REC reference: 16/NW/0507, IRAS project ID: 202688

Study design
Two-stage feasibility study with interventional, observational and cross-sectional components followed by the main study phase

Primary study design
Observational

Secondary study design
Epidemiological study

Study setting(s)
Community

Study type(s)
Other

Participant information sheet
Available on trial website

Health condition(s) or problem(s) studied
Nasal carriage of Staphylococcus aureus

Interventions
Current interventions as of 24/02/2023:

Phase 1 and 2 of the study will involve the recruitment of participants who have taken part in previous studies (INTERVAL: ISRCTN24760606) and have agreed to be invited to other studies organised by NHS Blood and Transplant and/or University of Cambridge and/or University of Oxford.

Phase 3 (main study) will, in addition to INTERVAL, include participants from the COMPARE study (ISRCTN90871183) and STRIDES BioResource study.

In phase 1: Participant acceptability of collecting two versus three swabs will be compared in approximately 250 participants (125 per arm). Prior to invitation, there will be a 1:1 random allocation to each of the swabbing frequencies; simple block randomisation stratified by age and gender will be used. The acceptability of each protocol will be based on participant response rates to the two- and three-swab protocol and participant feedback from an online questionnaire asking about experiences of taking part in the study – now complete.

Phase 2: A further (approximately) 2000 participants will be invited to collect nasal swabs. The number of swabs collected will be informed by the results of phase 1 (i.e. either two or three). This phase will assess, more generally, the operational feasibility of collecting, processing and analysing swabs in large numbers of participants. It will also assess lifestyle and biomarker correlates of persistent nasal S. aureus carriage. – now complete.

In Phase 3: This is the main study phase where we will consent enough participants to retrieve a full sample set (3 swabs) from 20,000 participants. . This phase will assess, more generally, the operational feasibility of collecting, processing and analysing swabs in large numbers of participants. It will also assess lifestyle and biomarker correlates of persistent nasal S. aureus carriage and to identify new genetic associations.

In all phases of the study prevalence rates of S. aureus carriage will be assessed.

Participants joining the study will be asked to complete an online baseline questionnaire designed to collect health and lifestyle information which may be relevant to predisposition to S. aureus carriage. On completion of their questionnaire, participants will be sent a nasal swabbing pack in the post and asked to return their first swab immediately and subsequent swabs at weekly intervals. Once participants have returned all swabs they will be sent an online feedback questionnaire about their experiences of taking the swabs and participating in the study.

The study will be administered by the University of Cambridge and will be conducted in the community across England.

_____

Previous interventions as of 19/06/2019:

Phase 1 and 2 of the study will involve the recruitment of participants who have taken part in previous studies (INTERVAL: ISRCTN24760606) and have agreed to be invited to other studies organised by NHS Blood and Transplant and/or University of Cambridge and/or University of Oxford.

Phase 2 (extended) (main study) will, in addition to INTERVAL, include participants from the COMPARE study (ISRCTN90871183).

In phase 1: Participant acceptability of collecting two versus three swabs will be compared in approximately 250 participants (125 per arm). Prior to invitation, there will be a 1:1 random allocation to each of the swabbing frequencies; simple block randomisation stratified by age and gender will be used. The acceptability of each protocol will be based on participant response rates to the two- and three-swab protocol and participant feedback from an online questionnaire asking about experiences of taking part in the study – now complete.

Phase 2: A further (approximately) 2000 participants will be invited to collect nasal swabs. The number of swabs collected will be informed by the results of phase 1 (i.e. either two or three). This phase will assess, more generally, the operational feasibility of collecting, processing and analysing swabs in large numbers of participants. It will also assess lifestyle and biomarker correlates of persistent nasal S. aureus carriage. – now complete.

In Phase 2 (extended): This is the main study phase where approximately 20,000 participants will be invited to take part. This phase will assess, more generally, the operational feasibility of collecting, processing and analysing swabs in large numbers of participants. It will also assess lifestyle and biomarker correlates of persistent nasal S. aureus carriage and to identify new genetic associations.

In all phases of the study prevalence rates of S. aureus carriage will be assessed.

Participants joining the study will be asked to complete an online baseline questionnaire designed to collect health and lifestyle information which may be relevant to predisposition to S. aureus carriage. On completion of their questionnaire, participants will be sent a nasal swabbing pack in the post and asked to return their first swab immediately and subsequent swabs at weekly intervals. Once participants have returned all swabs they will be sent an online feedback questionnaire about their experiences of taking the swabs and participating in the study.

The study will be administered by the University of Cambridge and will be conducted in the community across England.

_____

Previous interventions:

Both stages of the study will involve the recruitment of participants who have taken part in a previous study (INTERVAL: ISRCTN24760606) and have agreed to be invited to other studies

organised by NHS Blood and Transplant and/or University of Cambridge and/or University of Oxford.

In stage 1: Participant acceptability of collecting two versus three swabs will be compared in approximately 250 participants (125 per arm). Prior to invitation, there will be a 1:1 random allocation to each of the swabbing frequencies; simple block randomisation stratified by age and gender will be used. The acceptability of each protocol will be based on participant response rates to the two- and three-swab protocol and participant feedback from an online questionnaire asking about experiences of taking part in the study.

In Stage 2: A further (approximately) 2000 participants will be invited to collect nasal swabs. The number of swabs collected will be informed by the results of Stage 1 (i.e. either two or three). This stage will assess, more generally, the operational feasibility of collecting, processing and analysing swabs in large numbers of participants. It will also assess lifestyle and biomarker correlates of persistent nasal S. aureus carriage.

In both stages of the study prevalence rates of S. aureus carriage will be assessed.

Participants joining the study will be asked to complete an online baseline questionnaire designed to collect health and lifestyle information which may be relevant to predisposition to S. aureus carriage. On completion of their questionnaire, participants will be sent a nasal swabbing pack in the post and asked to return their first swab immediately and subsequent swabs at weekly intervals. Once participants have returned all swabs they will be sent an online feedback questionnaire about their experiences of taking the swabs and participating in the study.

The study will be administered by the University of Cambridge and will be conducted in the community across England.

Intervention Type
Other

Primary outcome measure
Current primary outcome measure as of 19/06/2019:

1. To assess the feasibility of collecting, processing and analysing nasal swabs in large numbers of participants.
2. To assess the prevalence of persistent S. aureus nasal carriage rates in healthy volunteers.
3. To assess lifestyle and biomarker correlates of persistent nasal S. aureus carriage and validate associations with previously suggested candidate genetic variants and to identify new genetic associations.

_____

Previous primary outcome measure:

Response rates to nasal swab collections including differences in response rates to initial invitation to provide two vs three swabs and compliance to the collection and return of two vs three swabs

Secondary outcome measures
Current secondary outcome measures as of 27/02/2023:

Phases 1 & 2:
1. Participant feedback from the online questionnaire administered to participants on completion of their involvement in the study. Data collected will include binary (yes/no), Likert Scale and free text responses
2. Operational outcomes: the success of operational procedures will be reviewed including: posting of nasal swabbing packs, sample receipt at the processing laboratory and laboratory methods for sample processing and analysis

Phase 3
1. To assess the feasibility of collecting, processing and analysing nasal swabs in large numbers of participants.
2. To assess the prevalence of persistent S. aureus nasal carriage rates in healthy volunteers.
3. To assess lifestyle and biomarker correlates of persistent nasal S. aureus carriage and validate associations with previously suggested candidate genetic variants and to identify new genetic associations.

_____

Previous secondary outcome measures as of 19/06/2019:

Phases 1 & 2:
1. Participant feedback from the online questionnaire administered to participants on completion of their involvement in the study. Data collected will include binary (yes/no), Likert Scale and free text responses
2. Operational outcomes: the success of operational procedures will be reviewed including: posting of nasal swabbing packs, sample receipt at the processing laboratory and laboratory methods for sample processing and analysis

_____

Previous secondary outcome measures:

Stages 1 & 2:
1. Participant feedback from the online questionnaire administered to participants on completion of their involvement in the study. Data collected will include binary (yes/no), Likert Scale and free text responses
2. Prevalence of persistent nasal carriage of Staphylococcus aureus in healthy volunteers. The point prevalence of carriage will be calculated using the first swabs returned by participants. Persistent carriage will be defined based on two positive swabs in participants randomised to the two-swab protocol and three positive swabs in participants randomised to the three-swab protocol.
3. Operational outcomes: the success of operational procedures will be reviewed including: posting of nasal swabbing packs, sample receipt at the processing laboratory and laboratory methods for sample processing and analysis

Stage 2:
Lifestyle, biomarker and genetic correlates of persistent S. aureus carriage assessed using either logistic or linear regression models

Overall study start date
01/09/2016

Completion date
27/06/2026

Eligibility

Key inclusion criteria
Current inclusion criteria as of 24/02/2023:

1. Be already enrolled in the INTERVAL, COMPARE or STRIDES BioResource studies
2. Have an email address for study participation
3. Reside in mainland England
4. Have a good understanding of the English language, both written and oral (study materials are not tailored to support non-English language speakers)

Previous participant inclusion criteria as of 19/06/2019:

1. Be already enrolled in the INTERVAL or COMPARE studies
2. Have an email address for study participation
3. Reside in mainland England
4. Have a good understanding of the English language, both written and oral (study materials are not tailored to support non-English language speakers)

Previous participant inclusion criteria:

1. Be already enrolled in the INTERVAL study
2. Have an email address for study participation
3. Reside in mainland England
4. Have a good understanding of the English language, both written and oral (study materials are not tailored to support non-English language speakers)

Participant type(s)
Healthy volunteer

Age group
Adult

Sex
Both

Target number of participants
Phase 1: 250, Phase 2: 2,000, Main phase: 20,000

Key exclusion criteria
Current exclusion criteria as of 24/02/2023:

1. Received three invitations from the INTERVAL, COMPARE or STRIDES BioResource research study teams to take part in other studies in the past year

2. Withdrawn their consent to take part in the INTERVAL,COMPARE or STRIDES BioResource studies
3. Any abnormalities or ongoing medical conditions that affect the nose (e.g. nosebleeds, nasal polyps, rhinitis, etc)
4. Undergone nasal surgery in the last year
5. Nasal fracture in the last year

_____

Previous participant exclusion criteria as of 19/06/2019:

1. Received three invitations from the INTERVAL or COMPARE research study teams to take part in other studies in the past year
2. Withdrawn their consent to take part in the INTERVAL or COMPARE studies
3. Any abnormalities or ongoing medical conditions that affect the nose (e.g. nosebleeds, nasal polyps, rhinitis, etc)
4. Undergone nasal surgery in the last year
5. Nasal fracture in the last year

_____

Previous participant exclusion criteria:

1. Have received 3 invitations from the INTERVAL research study team to take part in other studies in the past year
2. Have withdrawn their consent to take part in the INTERVAL study
3. Have any abnormalities or ongoing medical conditions that affect the nose (e.g. nosebleeds, nasal polyps, rhinitis, etc)
4. Have undergone nasal surgery in the last year
5. Have had a nasal fracture in the last year
6. Are not going to be available to take part in the study during the next two months due to reasons such as an extended holiday

Date of first enrolment
19/09/2016

Date of final enrolment
31/12/2023

Locations

Countries of recruitment
England

United Kingdom

Study participating centre
University of Cambridge
Department of Medicine

Box 157, level 5
Addenbrooke's Hospital
Hills Road
Cambridge
United Kingdom
CB2 0QQ

Study participating centre
University of Cambridge
Dept Public Health and Primary Care
Heart and Lung Research Institute
Papworth Road
Cambridge
United Kingdom
CB2 0BB

Sponsor information

Organisation
University of Cambridge (UK)

Sponsor details
School of Clinical Medicine
Addenbrooke's Hospital Hills Road
Cambridge
England
United Kingdom
CB2 0SP

Sponsor type
University/education

ROR
https://ror.org/013meh722

Funder(s)

Funder type
Charity

Funder Name
Wellcome Trust

Alternative Name(s)

Funding Body Type
Private sector organisation

Funding Body Subtype
International organizations

Location
United Kingdom

Funder Name
UK Research and Innovation

Alternative Name(s)
UKRI

Funding Body Type
Government organisation

Funding Body Subtype
National government

Location
United Kingdom

Funder Name
Health Data Research (HDR) UK

Funder Name
Isaac Newton Trust

Alternative Name(s)

Funding Body Type
Private sector organisation

Funding Body Subtype
Universities (academic only)

Location
United Kingdom

Results and Publications

Publication and dissemination plan

The findings from this study will be written up for publication in scientific journals and presentation to other researchers. In due course we will also provide details of the findings of the study on the CARRIAGE website.

Anonymous information and samples may be made available on request to researchers who have relevant scientific and ethics approvals for their planned research.

Intention to publish date

31/12/2024

Individual participant data (IPD) sharing plan

Not provided at time of registration

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