



PRINCIPLE: Platform Randomised trial of INTERventions against COVID-19 In older people

Chief Investigator: Professor Chris Butler

Tel: +44 (0)1865 289670

Email: christopher.butler@phc.ox.ac.uk

Address: Primary Care Clinical Trials Unit

Gibson Building, 1st Floor, Radcliffe Observatory Quarter, Woodstock Road, Oxford. OX2 6GG

Date:

REC Substantial Amendment 8

IRAS ID: 281958

Dear REC committee members,

Subject: REC Substantial Amendment 8 for PRINCIPLE Trial in relation to COVID-19

Please find attached the Amendment submission package for the trial entitled: Platform Randomised trial of INTERventions against COVID-19 In older people (PRINCIPLE). The amendments are described and justified below.

Certain documents have been updated or created for the trial.

Summary of key proposed changes to PRINCIPLE Trial: Amendment 8

1. **Translated video script** has been created for local and national promotion of the trial. The video will feature on the trial website and will also be translated into other languages with the aim of reaching BAME communities.
2. **Addition of a non-NHS site as a recruitment site**
BrisDoc Healthcare Services Ltd
3. **Text Message invite has been simplified** to update wording around the test result into the lay terminology, this has previously caused confusion for patients.
4. **Video infographic script** has been created for local and national promotion of the trial. The infographic will provide a clear summary of the trial
5. **Patient recruitment poster specifically for use in pharmacies** has been created, for promotion of the trial in pharmacy settings.
6. **End of follow up letter** has been updated to include a link to the NIHR Participant Research Experience Survey. The information collected could provide important learnings for other centralised studies in the future. It is stated that this is optional.

Yours sincerely,

Dr Hannah Swayze

On behalf of Professor Christopher Butler

Table 1: List of documentation submitted to REC (new documents highlighted in blue)

Document	Version	Date	Key point (described above) documents relate to
Pictorial Translated Video Script	1.0	12.10.2020	1
PRINCIPLE Text Message Info (tracked changes & clean)	4.1	12.10.2020	3
PRINCIPLE Video Infographic Script	1.0	12.10.2020	4
PRINCIPLE Patient Recruitment Poster Pharmacy	1.0	12.10.2020	5
PRINCIPLE End of Follow Up Letter (tracked changes & clean)	1.1	12.10.2020	6
Non-NHS-HSC Site Assessment Form (BrisDoc)	1.1	12.10.2020	2
CV of PI for Non-NHS Site	1.0		2
Locked Amendment Tool	SA8	12.10.2020	All

PRINCIPLE TRIAL: TEXT MESSAGE INFORMATION FOR PARTICIPANTS

This is a message from Dr XX at XX Medical Practice. If you experience **symptoms of COVID-19** and have had them for **less than 14 days and are not starting to feel better OR** you have had a **positive test** for COVID-19, AND are unwell you may be eligible to join a clinical trial: www.principletrial.org to find out more or call the Trial Team: 0800 138 0880

PRINCIPLE TRIAL: TEXT MESSAGE INFORMATION FOR PARTICIPANTS (SHORT VERSION 1)

This is from Dr xxxxxxxx at XX. If you currently feel unwell with COVID-19 symptoms **OR** you have had a **positive test** for COVID-19 infection AND are unwell, and would like to know about joining a clinical trial click www.principletrial.org or call 0800 138 0880

PRINCIPLE video infographic script

Introduction

COVID-19 can lead to significant medical problems, hospitalisation, and sometimes death. The risk of complications from COVID-19 is generally greater in people aged 50 years and older with existing health conditions and in those aged 65 years and older. So far, there are no treatments that have been proven in clinical trials to help people recover from COVID-19 while in the community.

We need to find treatments that help people recover quicker and reduce the need for them to go to hospital.

What the study involves

We are looking for people aged 50 years or over with symptoms including a cough OR temperature OR loss/change of smell/taste, starting within the past 14 days. OR you may have had a recent confirmed test for COVID-19 with any other symptoms.

Registering for the study can be completed online or over the phone and no face to face visits are required. You will be randomly assigned to receive usual care or medication. We will send everyone a participant pack containing a testing swab and medication if you are assigned to receive it, directly to their home. If you have any questions at all you can contact the trial free phone number or your GP.

Summary

If you are unwell now with COVID-19 or **become unwell in the future**, please visit our website www.principletrial.org for more information or to register for the trial. You can also call the team on 0800 138 0880

Please help us fight COVID-19 and consider registering for the PRINCIPLE Trial.



Researchers at the University of Oxford are appealing for people with typical COVID-19 symptoms, aged 50-64 years with existing medical conditions, or 65 years and over with or without existing medical conditions, to take part in a COVID-19 treatment trial - PRINCIPLE.

Currently, there are no effective treatments available that have been shown through clinical trials to reduce disease burden in the community. PRINCIPLE is evaluating whether certain commonly used medicines, such as antibiotics, may prevent patients with COVID-19 from becoming more unwell and needing hospital care.

The research team are appealing for people experiencing symptoms that are likely to be caused by a COVID-19 infection, for fewer than 15 days, to join the trial. People may also be eligible to join if they have had a positive test for COVID infection which was taken less than 15 days ago, and are unwell with any symptoms.

There are no face to face visits involved and people can take part either online or over the phone. Anyone in the UK over 50 who is unwell with COVID symptoms may be able to take part and a participant pack will be couriered directly to patient's homes.

<Quote from healthcare provider making video>

Help in the fight against COVID-19

The PRINCIPLE trial aims to find treatments that improve symptoms and reduce hospital admission for people with COVID-19

Anyone suspected or confirmed with COVID-19 may be eligible to join the PRINCIPLE Trial.

(Symptoms to look out for include *New Cough *High Temperature *Loss/change in smell/taste)

Open to anyone aged 65 and above?
Or aged 50 to 64 with any of these illnesses?



High blood pressure and/ or heart disease



Diabetes *not* treated with insulin



Obesity or BMI ≥ 35 kg/m²



Asthma or lung disease



Stroke or neurological problems



Weakened immune system due to serious illness or medication (e.g. chemotherapy).



Liver disease

To find out more or register for the study, please visit:

www.principletrial.org

Tel: 0800 138 0880

email: principle@phc.ox.ac.uk

IRAS PROJECT ID 281958, REC Reference 20/SC/0158 Confirmation of favourable opinion for substantial amendment

berkshire.rec@hra.nhs.uk <noreply@harp.org.uk>

Tue 10/13/2020 16:12

To: Christopher Butler <christopher.butler@phc.ox.ac.uk>; CTRG Sponsorship Correspondence <ctr@admin.ox.ac.uk>

Cc: nhsg.nrspcc@nhs.net <nhsg.nrspcc@nhs.net>; Hannah Swayze <hannah.swayze@phc.ox.ac.uk>; research.amendments@hscni.net <research.amendments@hscni.net>

1 attachments (115 KB)

IRAS 281958 SL32_Favourable_opinion_of_a_substantial_amendment.pdf;

Dear Professor Butler

IRAS project ID:	281958
REC reference:	20/SC/0158
Short Study title:	PRINCIPLE [COVID-19] [UPH]
Date complete amendment submission received:	12 October 2020
Amendment No./ Sponsor Ref:	Substantial Amendment 8
Amendment Date:	12 October 2020
Amendment Type:	Substantial
Outcome of HRA Assessment	This email also constitutes HRA and HCRW Approval for the amendment, and you should not expect anything further.

I am pleased to confirm that this amendment has been reviewed by the Research Ethics Committee and has received a Favourable Opinion. Please find attached a copy of the Favourable Opinion letter.

HRA and HCRW Approval Status

As detailed above, **this email also constitutes HRA and HCRW Approval for the amendment.** No separate confirmation of HRA and HCRW Approval will be issued.

User Feedback

The Health Research Authority is continually striving to provide a high quality service to all applicants and sponsors. You are invited to give your view of the service you have received and the application procedure. If you wish to make your views known please use the feedback form available on the HRA website: <http://www.hra.nhs.uk/about-the-hra/governance/quality-assurance/>.

If you require further information, please contact me.

Kind regards

Alison Doherty
Approvals Administrator

Bristol REC Centre | Whitefriars | BS1 2NT

T. 020 7104 8049

E. berkshire.rec@hra.nhs.uk

W. www.hra.nhs.uk

Sign up to receive our newsletter [HRA Latest](#).

South Central - Berkshire Research Ethics Committee

Bristol REC Centre
Whitefriars
Level 3, Block B
Lewins Mead
Bristol
BS1 2NT

13 October 2020

Professor Butler
University of Oxford, Joint Research Office
1st floor, Boundary Brook House, Churchill Drive, Headington
Oxford
OX3 7GB

Dear Professor Butler

Study title: Platform Randomised trial of INterventions against COVID-19 In older peoPLE
REC reference: 20/SC/0158
Protocol number: PRINCIPLE
EudraCT number: 2020-001209-22
Amendment number: Substantial Amendment 8
Amendment date: 12 October 2020
IRAS project ID: 281958

The above amendment was reviewed by the Sub-Committee in correspondence.

Ethical opinion

The members of the Committee taking part in the review gave a favourable ethical opinion of the amendment on the basis described in the notice of amendment form and supporting documentation.

Approved documents

The documents reviewed and approved at the meeting were:

<i>Document</i>	<i>Version</i>	<i>Date</i>
Completed Amendment Tool [SA8 Locked Amendment Tool]	n/a	12 October 2020
Copies of advertisement materials for research participants [Patient Recruitment Pharmacy Poster]	1.0	12 October 2020
Cover Letter [REC SA8 Cover Letter]	n/a	12 October 2020
Non-NHS/HSC Site Assessment Form [Non-NHS Assessment Form]	1.1	12 October 2020
Other [CV for PI of Non-NHS]	1.0	24 September 2020
Other [Text Message Info Clean]	4.1	12 October 2020
Other [Text Message Info Tracked changes]	4.1	12 October 2020

Other [Video Infographic Script]	1.0	12 October 2020
Other [End of Follow up letter Clean]	1.1	12 October 2020
Other [End of Follow up letter tracked changes]	1.1	12 October 2020
Other [Translated video script]	1.0	12 October 2020

Membership of the Committee

The members of the Committee who took part in the review are listed on the attached sheet.

Working with NHS Care Organisations

Sponsors should ensure that they notify the R&D office for the relevant NHS care organisation of this amendment in line with the terms detailed in the categorisation email issued by the lead nation for the study.

Amendments related to COVID-19

We will update your research summary for the above study on the research summaries section of our website. During this public health emergency, it is vital that everyone can promptly identify all relevant research related to COVID-19 that is taking place globally. If you have not already done so, please register your study on a public registry as soon as possible and provide the HRA with the registration detail, which will be posted alongside other information relating to your project.

Statement of compliance

This Committee is recognised by the United Kingdom Ethics Committee Authority under the Medicines for Human Use (Clinical Trials) Regulations 2004, and is authorised to carry out the ethical review of clinical trials of investigational medicinal products.

The Committee is fully compliant with the Regulations as they relate to ethics committees and the conditions and principles of good clinical practice.

The Committee is constituted in accordance with the Governance Arrangements for Research Ethics Committees and complies fully with the Standard Operating Procedures for Research Ethics Committees in the UK.

HRA Learning

We are pleased to welcome researchers and research staff to our HRA Learning Events and online learning opportunities– see details at: <https://www.hra.nhs.uk/planning-and-improving-research/learning/>

IRAS Project ID - 281958:

Please quote this number on all correspondence

Yours sincerely
PP



Mr David Carpenter
Chair

E-mail: berkshire.rec@hra.nhs.uk

Enclosures: List of names and professions of members who took part in the review

Copy to: Christopher Butler, University of Oxford

South Central - Berkshire Research Ethics Committee

Attendance at Sub-Committee of the REC meeting on 23 October 2020

Committee Members:

<i>Name</i>	<i>Profession</i>	<i>Present</i>	<i>Notes</i>
Mr David Carpenter	Retired Social Scientist	Yes	
Dr Mike Proven	University Research Governance Officer	Yes	

Also in attendance:

<i>Name</i>	<i>Position (or reason for attending)</i>
Miss Alison Doherty	Approvals Administrator