# The AVICCI study: a feasibility study to assess the effects of AntiretroViral Intensification with Cenicriviroc for the management of HIV-associated Cognitive Impairment

<b>Submission date</b> 09/10/2015	<b>Recruitment status</b> No longer recruiting	Prospectively registered
		☐ Protocol
Registration date	Overall study status	Statistical analysis plan
03/03/2016	Completed	[X] Results
<b>Last Edited</b> 21/06/2019	Condition category Infections and Infestations	[] Individual participant data

#### Plain English summary of protocol

Background and study aims

HIV is a virus that attacks the immune system, weakening the body's ability to fight infections and disease. HIV is treated with a combination of antiretroviral drugs which stops the virus replicating in the body and allows your immune system to recover. This prevents you from developing many illnesses related to the HIV virus. However, some people can still develop attention and memory problems known as cognitive function impairment. The aim of this study is to see if adding in a new antiretroviral drug has an effect on cognitive function.

#### Who can participate?

Patients aged 18 or over with HIV and clinically significant cognitive impairment.

#### What does the study involve?

The study is up to 16 weeks long and there are a total of 6 visits. For 8 weeks participants receive the new medication in addition to their current treatment. At the start and end of the study participants have a lumbar puncture (where a needle is inserted into the lower part of the spine) and an MRI scan and on three occasions complete a series of tests to assess their brain function. There are safety blood tests at all visits.

#### What are the possible benefits and risks of participating?

There is no direct benefit to the participants but the information gained from this study may be of benefit in the treatment of HIV-infected patients in the future. There are possible side effects of the treatment that include diarrhoea, vomiting, rash, headache and tiredness. There are also risks associated with having a lumbar puncture, including pain at the site and headaches.

Where is the study run from? Imperial College Healthcare NHS Trust (UK)

When is the study starting and how long is it expected to run for? October 2015 to September 2016

Who is funding the study? National Institute for Health Research (UK)

Who is the main contact? Mr Kenneth Legg k.legg@imperial.ac.uk

# **Contact information**

#### Type(s)

Public

#### Contact name

Mr Kenneth Legg

#### Contact details

Clinical Trials Centre St Mary's Hospital London United Kingdom W2 1NY +44 (0)20 3312 1464 k.legg@imperial.ac.uk

## Type(s)

Scientific

#### Contact name

Dr Alan Winston

#### **ORCID ID**

http://orcid.org/0000-0002-5847-8416

#### Contact details

Clinical Trials Centre Winston Churchill Wing St. Mary's Hospital Praed Street London United Kingdom W2 1NY

\_

a.winston@imperial.ac.uk

# Additional identifiers

**EudraCT/CTIS** number

#### **IRAS** number

#### ClinicalTrials.gov number

# Secondary identifying numbers

AVICCI01

# Study information

#### Scientific Title

A feasibility study to assess the effects of AntiretroViral Intensification with Cenicriviroc for the management of HIV-associated Cognitive Impairment

#### **Acronym**

**AVICCI** 

#### Study objectives

To assess if the addition of Cenicriviroc to current antiretroviral treatment is acceptable, safe and tolerable for persons living with HIV, to measure the amount of cenicriviroc in the cerebrospinal fluid, and to find out whether it improves cognitive function in patients previously identified as having clinical cognitive dysfunction.

#### Ethics approval required

Old ethics approval format

#### Ethics approval(s)

Brighton & Sussex REC South-East Coast, approved 24/11/2015, amendment 1 05/01/2016, amendment 2 25/01/2016, ethics ref: 15-LOC-1887

#### Study design

Phase II feasibility study

# Primary study design

Interventional

# Secondary study design

Non randomised study

# Study setting(s)

Hospital

## Study type(s)

Treatment

# Participant information sheet

Not available in web format, please use the contact details below to request a patient information sheet

#### Health condition(s) or problem(s) studied

HIV

#### **Interventions**

This is a phase II feasibility study to assess changes in patients who have been identified as having cognitive function impairment following the addition of a new drug to an existing effective regime of three HIV antiretroviral drugs.

This is an intensification study of patients stable on a triple combination of HIV antiretroviral therapy. Cenicriviroc at a dose of either 25 mg, 150 mg or 300 mg as oral tablets will be taken once daily (dose is dependent on the participants current treatment regime). The Cenicriviroc will be taken for 8 weeks and a safety visit will be performed 4 weeks after discontinuing the trial drug. This is a single arm study.

#### Intervention Type

Drug

#### Phase

Phase II

#### Drug/device/biological/vaccine name(s)

Cenicriviroc

#### Primary outcome measure

- 1. Acceptability (measured via patient questionnaires) measured at week 8
- 2. Pharmacokinetic results (measured via CSF and plasma pharmacokinetics) measured at week 8
- 3. Cerebral function parameters (cognitive testing, neuro-imaging modalities) measured at week 8

#### Secondary outcome measures

Descriptive results of patient questionnaires and laboratory parameters measured at week 8

#### Overall study start date

01/10/2015

#### Completion date

30/09/2016

# **Eligibility**

#### Key inclusion criteria

- 1. Aged 18 or over at screening, male or female subjects
- 2. Documented HIV-infected
- 3. Undetectable plasma HIV RNA (<200 copies/mL) for at least 6 months
- 4. Demonstrated clinically significant cognitive impairment
- 5. On cART comprising of BHIVA guideline recommended therapies (2015 guidelines) with the exception of elvitegravir/cobicistat and rilpivirine
- 6. Comorbidities, if present, are stably managed for at least 6 months
- 7. No clinically significant recreational drug use or alcohol dependence
- 8. Male subjects who are heterosexually active must use two forms of barrier contraception (e.

- g., condom and diaphragm) during heterosexual intercourse, from screening through completion of the study
- 9. Female subjects may be eligible to enter and participate into the study if she:
- 9.1. Is of non-child-bearing potential defined as either post-menopausal (12 months of spontaneous amenorrhea and ≥ 45 years of age) or physically incapable of becoming pregnant with documented tubal ligation, hysterectomy or bilateral oophorectomy
- 9.2. Is of child-bearing potential with a negative pregnancy test at both Screening and Day 0 and agrees to use one of the following methods of contraception to avoid pregnancy:
- 9.2.1. Complete abstinence from penile-vaginal intercourse from 2 weeks prior to administration of study drug, throughout the study, and for at least 2 weeks after discontinuation of all study medications
- 9.2.2. Double barrier method (male condom/spermicide, male condom/diaphragm, diaphragm/spermicide)
- 9.2.3. Any intrauterine device (IUD) with published data showing that the expected failure rate is <1% per year (not all IUDs meet this criterion)
- 9.2.4. Male partner sterilization confirmed prior to the female subject's entry into the study, and this male is the sole partner for that subject
- 9.2.5. Approved hormonal contraception
- 9.2.6. Any other method with published data showing that the expected failure rate is <1% per year

Any contraception method must be used consistently, in accordance with the approved product label and for at least 2 weeks after discontinuation of study medication.

#### Participant type(s)

Patient

#### Age group

Adult

#### Lower age limit

18 Years

#### Sex

Both

# Target number of participants

10

#### Total final enrolment

7

#### Key exclusion criteria

- 1. Current major depression (score of less than 15 on PHQ-9 score at study screening)
- 2. Chronic neurological diseases (e.g., epilepsy and stroke; at the discretion of the investigator)
- 3. History of severe head injury (with loss of consciousness for >30 minutes)
- 4. Cerebral AIDS defining infections
- 5. Current intravenous drug use (past six months)
- 6. Severe psychiatric disease (at the discretion of the investigator)
- 7. Contra-indication for MRI scan (e.g., claustrophobia, metal in body, physically unable to lie flat)
- 8. Contraindications to lumbar-puncture examination (at the discretion of the investigator)
- 9. Current or previous use of CCR5 inhibitors (maraviroc, cenicriviroc or others)

- 10. Disallowed medication which may interact with cenicriviroc
- 11. Chronic liver disease
- 12. Laboratory investigations at screening out with the follow cut-offs:
- 12.1. Haemoglobin < 8.5 g/dL
- 12.2. Absolute neutrophil count < 1000
- 12.3. Platelet count < 100,000
- 12.4. ALT or AST> 5 x upper limit of normal
- 12.5. Estimated creatinine clearance < 50 mL/min (Cockcroft and Gault 1979)
- 13. In the opinion of the investigator unable to comply with study procedures

#### Date of first enrolment

01/10/2015

#### Date of final enrolment

30/11/2015

# Locations

#### Countries of recruitment

England

**United Kingdom** 

# Study participating centre Imperial College Healthcare NHS Trust

Clinical Trials Centre
Winston Churchill Wing
St Mary's Hospital
Praed Street
London
United Kingdom
W2 1NY

# Sponsor information

#### Organisation

Imperial College London

#### Sponsor details

Joint Research Compliance Office Imperial College London and Imperial College Healthcare NHS Trust Room 5L10C, 5th Floor, Lab Block Charing Cross Hospital Fulham Palace Road London England United Kingdom W6 8RF

#### Sponsor type

University/education

#### **ROR**

https://ror.org/041kmwe10

# Funder(s)

#### Funder type

Government

#### **Funder Name**

National Institute for Health Research

#### Alternative Name(s)

National Institute for Health Research, NIHR Research, NIHRresearch, NIHR - National Institute for Health Research, NIHR (The National Institute for Health and Care Research), NIHR

#### **Funding Body Type**

Government organisation

#### **Funding Body Subtype**

National government

#### Location

**United Kingdom** 

# **Results and Publications**

# Publication and dissemination plan

To be confirmed at a later date

Intention to publish date

Individual participant data (IPD) sharing plan

#### IPD sharing plan summary

Available on request

#### **Study outputs**

Output type Details Date created Date added Peer reviewed? Patient-facing?