

A trial of rivastigmine to prevent falls in Parkinson's

Submission date 08/04/2019	Recruitment status No longer recruiting	<input checked="" type="checkbox"/> Prospectively registered <input checked="" type="checkbox"/> Protocol
Registration date 16/04/2019	Overall study status Completed	<input type="checkbox"/> Statistical analysis plan <input type="checkbox"/> Results
Last Edited 23/08/2024	Condition category Nervous System Diseases	<input type="checkbox"/> Individual participant data <input checked="" type="checkbox"/> Record updated in last year

Plain English summary of protocol

Background and study aims

Parkinson's disease is a common condition particularly affecting older people. Falls are a very frequent complication of the disease affecting 60% of people with Parkinson's every year. As the population ages, the number of people living with Parkinson's disease and the occurrence of complications will increase. The loss of the chemical dopamine in the brain causes walking in Parkinson's to become slower, unsteady and irregular. People with the condition are therefore at a very high risk of falling. To some extent, people can compensate for these changes by paying more attention to their walking. However, Parkinson's also diminishes memory and thinking ability. This decreases people's ability to pay attention to their walking, especially when doing something at the same time.

Who can participate?

Cholinesterase inhibitor (ChEis) are drugs that are currently used to treat people with memory problems in Parkinson's. The effect of these drugs on falls in Parkinson's has been tested to show that treatment has the potential to almost halve the number of falls.

This trial aims to definitively determine whether cholinesterase inhibitors (ChEi), can prevent falls in Parkinson's and whether this treatment is cost-effective.

What does the study involve?

600 participants with Parkinson's disease will be enrolled from hospitals throughout the UK. Participants will be randomly assigned to either receive the drug (ChEi) via a patch or receive a placebo (dummy) treatment via a patch. Neither the researchers nor the participants will know which group they are in. Participants will take the medication for 12 months and record any falls that they experience in diaries.

What are the possible benefits and risks of participating?

There are a few possible benefits of taking part in this trial. If allocated to the group that receives the active medication, walking unsteadiness and/or balance may improve and falls may be less likely, but there is no guarantee. The information we get from this study will improve the treatment of people with Parkinson's disease in the future.

There are also some risks and discomforts of the trial. The assessments are quite detailed and therefore may cause tiredness and fatigue.

Like all medicines, the treatment can cause side effects, although not everybody gets them. Because the medication is already used to treat memory problems we know a lot about the side effects.

Side effects are experienced more frequently when you start your medicine or increase the dose. In most cases, side effects will gradually disappear.

Where is the study run from?

Bristol Medical School, University of Bristol, UK

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

April 2018 to July 2024

Who is funding the study?

NIHR Evaluation, Trials and Studies Co-ordinating Centre (NETSCC)

Who is the main contact?

CHIEF-PD Trial Management Team, Chief-pd@bristol.ac.uk

Study website

<https://chiefpd.blogs.bristol.ac.uk/>

Contact information

Type(s)

Scientific

Contact name

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University of Bristol

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United Kingdom

BS8 1NU

+44 (0)117 4283111

Chief-pd@bristol.ac.uk

Additional identifiers

EudraCT/CTIS number

2018-003219-23

IRAS number

235625

ClinicalTrials.gov number

NCT04226248

Secondary identifying numbers

CPMS 40906, IRAS 235625

Study information

Scientific Title

CHIEF-PD (CHolinesterase Inhibitor to prEvent Falls in Parkinson's Disease): A phase 3 randomised double-blind placebo-controlled trial of rivastigmine to prevent falls in Parkinson's disease

Acronym

CHIEF-PD

Study objectives

Cholinesterase inhibitor (ChEi) treatment prevents people with Parkinson's from falling and is cost-effective.

Ethics approval required

Ethics approval required

Ethics approval(s)

Approved 17/05/2019, Southwest Central Bristol Research Ethics Committee (Health Research Authority, Whitefriars, Level 3 Block B, Lewins Mead, Bristol, BS1 2NT, United Kingdom; +44 (0) 2071048046; nrescommittee.southwest-bristol@nhs.net), ref: 19/SW/0043

Study design

Randomized controlled trial

Primary study design

Interventional

Secondary study design

Randomised controlled trial

Study setting(s)

Hospital

Study type(s)

Treatment

Participant information sheet

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

Health condition(s) or problem(s) studied

Parkinson's Disease

Interventions

Current interventions as of 05/12/2019:

Previous interventions:

600 participants with Parkinson's disease will be enrolled from hospitals throughout the UK. Participants will be randomly assigned to either receive the drug (ChEi) via a patch or receive a placebo (dummy) treatment via a patch. Participants are allocated to the two treatment arms using a technique that minimises the differences between the groups but still allocated people randomly. Neither the researchers nor the participants will know which group they are in. Participants will take the medication for 12 months and record any falls that they experience in diaries and through monthly telephone calls.

TRIAL ASSESSMENTS OVERVIEW

At baseline (enrolment) and follow-up (12 months) detailed tests of Parkinson's disease are undertaken to determine what effect the treatment has on the motor and non-motor symptoms of the condition. At monthly intervals between visits participants will complete a diary (detailed below).

BASELINE VISIT (2.5 hours)

Eligibility confirmation (10 minutes)

Consent (15 minutes)

Disease and functional measures (approx 1.5 hours)

1. Questionnaires (optionally can be completed pre-visit) (15 minutes):

1.1 Freezing of gait (New Freezing of Gait questionnaire NFOG-Q)

1.2 Fear of falling (Iconographical Falls Efficacy Scale [ICON-FES])

1.3 Mood (Geriatric Depression Scale [GDS])

1.4 Capability of older people (ICEpop CAPability measure for Older people [ICECAP-O])

1.5 Quality of life (EQ-5D-5L)

1.6 Starkstein Apathy Scale (SAS)

1.7 Swallowing (Swallowing Disturbance Questionnaire [SDQ])

2. Sociodemographics, medical and drug history

3. Brief examination: heart rate (+/- electrocardiogram if indicated), blood pressure, height and weight, Movement Disorder Society Unified Parkinson's Disease Scale (MDS-UPDRS) parts 3 and 4, frailty and gait assessment, Short Physical Performance Battery (SPPB).

4. Functional measures:

4.1 Cognition (Montreal Cognitive Assessment [MoCA])

SELF COMPLETION OF MONTHLY DIARY and PHONE CALLS

Falls are recorded in a patient diary which is posted back to the central team on a monthly basis. Participants are telephoned to remind them about the diaries, to identify any side effects that have occurred.

Questionnaire assessments of quality of life and healthcare use are completed in the diary at months 1, 3, 6, 9 and 12 in order to determine whether the treatment, if successful, is also cost-effective.

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Intervention Type

Drug

Phase

Phase III

Drug/device/biological/vaccine name(s)

Rivastigmine

Primary outcome measure

Fall rate measured using monthly diaries and telephone calls prospectively over 12 months from the day the IMP is commenced. A fall is defined as "unintentionally coming to rest on the ground or other lower surface without overwhelming external force or a major internal event"

Secondary outcome measures

Current secondary outcome measures as of 05/12/2019:

1. Parkinson's Disease markers assessed via the MDS-UPDRS total score in the practically defined ON state and each individual subscale (1-4) measured at baseline and 12 months

2. Freezing of gait assessed via the New Freezing Of Gait Questionnaire (NFOGQ) measured at

baseline and 12months

3. Frailty and physical performance assessed via Short Physical Performance Battery (SPPB), gait speed and frailty status measured at baseline and 12months
4. Cognition assessed via Montreal Cognitive Assessment (MoCA) measured at baseline and 12months
5. Depression assessed via Geriatric Depression Scale (GDS) measured at baseline and 12months
6. Apathy assessed via the Starkstein Apathy Scale (SAS)
7. Fear of falling assessed via the Iconographical Fall Efficacy Scale (ICON-FES) measured at baseline and 12months
8. Dysphagia assessed via the Swallowing Disturbance Questionnaire (SDQ) measured at baseline and 12months
9. Participant health-related quality of life assessed via the EuroQoL 5D-5L health status questionnaire (EQ-5D-5L) measured at baseline, 1 month, 3 months, 6 months, 9 months and 12months
10. Capability of older people assessed via the ICEpop CAPability measure for Older people (ICECAP-O) measured at baseline and 12 months
11. Mortality (all cause and PD related) through Office of National Statistics measured at 12months
12. Cost-effectiveness and NHS resource use through EQ-5D-5L and NHS Episode Statistics (HES) data

Secondary outcome via CHIEF-PD carer study:

12. Care-related quality of life via the Carer Experience Scale (CES) measured at baseline and 12months

Previous secondary outcome measures:

1. Parkinson's Disease markers assessed via the MDS-UPDRS total score in the practically defined ON state and each individual subscale (1-4) measured at baseline and 12months
2. Freezing of gait assessed via the New Freezing Of Gait Questionnaire (NFOGQ) measured at baseline and 12months
3. Frailty and physical performance assessed via Short Physical Performance Battery (SPPB), gait speed and frailty status measured at baseline and 12months
4. Cognition assessed via Montreal Cognitive Assessment (MoCA) measured at baseline and 12months
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6. Fear of falling assessed via the Iconographical Fall Efficacy Scale (ICON-FES) measured at baseline and 12months
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Secondary outcome via CHIEF-PD carer study:

12. Care-related quality of life via the Carer Experience Scale (CES) measured at baseline and 12months

Overall study start date

01/04/2018

Completion date

30/07/2024

Eligibility

Key inclusion criteria

Current participant inclusion criteria as of 05/07/2019:

1. Diagnosis of idiopathic Parkinson's disease
2. Modified Hoehn and Yahr stage 1-4 disease
3. Have experienced a fall in the previous year
4. Able to walk ≥ 10 m without aids or assistance
5. 18 years of age or above

Previous participant inclusion criteria:

1. Diagnosis of idiopathic Parkinson's disease
2. Modified Hoehn and Yahr stage 1-4 disease
3. Have experienced a fall in the previous year
4. Able to walk ≥ 10 m without aids or assistance

Participant type(s)

Patient

Age group

Adult

Lower age limit

18 Years

Sex

Both

Target number of participants

Planned Sample Size: 600; UK Sample Size: 600

Total final enrolment

600

Key exclusion criteria

Current participant exclusion criteria as of 05/07/2019:

1. Previous ChEi use in 12 months prior to enrolment.
2. Hypersensitivity to rivastigmine
3. Dementia diagnosed according to Movement Disorder Society (MDS) criteria
4. Inability to attend or comply with treatment or follow-up scheduling
5. Non-English-speaking patients (cognitive tests performed in English)
6. Falling ≥ 4 x per day

7. Unwillingness to use an acceptable method of contraception for the duration of the trial if they are of childbearing potential
8. Pregnant or breast feeding

Previous participant exclusion criteria:

1. Previous ChEi use in 12 months prior to enrolment
2. Hypersensitivity to rivastigmine
3. Dementia diagnosed according to Movement Disorder Society (MDS) criteria
4. Inability to attend or comply with treatment or follow-up scheduling
5. Non-English-speaking patients (cognitive tests performed in English)
6. Falling ≥ 4 x per day
7. Unwillingness to use an acceptable method of contraception for the duration of the trial if they are of childbearing potential

Date of first enrolment

02/09/2019

Date of final enrolment

30/04/2023

Locations

Countries of recruitment

England

Scotland

United Kingdom

Wales

Study participating centre

Royal United Hospitals Bath NHS Foundation Trust

Combe Park

Bath

United Kingdom

BA1 3NG

Study participating centre

Salford Royal NHS Foundation Trust

Stott Lane

Salford

United Kingdom

M6 8HD

Study participating centre

Northern Care Alliance

North Manchester General Hospital
Delaunays Road
Crumpsall
Manchester
United Kingdom
M8 5RB

Study participating centre

Royal Cornwall Hospitals NHS Trust

Royal Cornwall Hospital
Treliske
Truro
United Kingdom
TR1 3LJ

Study participating centre

NHS Lothian

Waverley Gate
2-4 Waterloo Place
Edinburgh
United Kingdom
EH1 3EG

Study participating centre

Derby Teaching Hospitals NHS Foundation Trust

Royal Derby Hospital
Uttoxeter Road
Derby
United Kingdom
DE22 3NE

Study participating centre

Plymouth Hospitals NHS Trust

Derriford Hospital
Derriford Road
Plymouth
United Kingdom
PL6 8DH

Study participating centre
Imperial College Healthcare NHS Trust
St. Marys Hospital
Praed Street
London
United Kingdom
W2 1NY

Study participating centre
Yeovil District Hospital NHS Foundation Trust
Yeovil District Hospital
Higher Kingston
Yeovil
United Kingdom
BA21 4AT

Study participating centre
The Newcastle Upon Tyne Hospitals NHS Foundation Trust
Freeman Hospital
Freeman Road
High Heaton
Newcastle-upon-Tyne
United Kingdom
NE7 7DN

Study participating centre
Taunton And Somerset NHS Foundation Trust
Musgrove Park Hospital
Taunton
United Kingdom
TA1 5DA

Study participating centre
Cambridge University Hospitals NHS Foundation Trust
Addenbrookes Hospital
Hills Road
Cambridge
United Kingdom
CB2 0QQ

Study participating centre

Royal Devon And Exeter NHS Foundation Trust

Royal Devon & Exeter Hospital
Barrack Road
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United Kingdom
EX2 5DW

Study participating centre

The Walton Centre NHS Foundation Trust

Lower Lane
Liverpool
United Kingdom
L9 7LJ

Study participating centre

University College London Hospitals NHS Foundation Trust

250 Euston Road
London
United Kingdom
NW1 2PG

Study participating centre

Barking, Havering and Redbridge University Hospitals NHS Trust Rom Valley Way, Romford, RM7 0AG

Rom Valley Way
Romford
United Kingdom
RM7 0AG

Study participating centre

Gateshead NHS Foundation Trust

Queen Elizabeth Avenue
Gateshead
United Kingdom
NE9 6SX

Study participating centre

Great Western Hospitals NHS Foundation Trust

Marlborough Road

Swindon
United Kingdom
SN3 6BB

Study participating centre

NHS Tayside
230 Clepington Road
Dundee
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Study participating centre

Nottingham University Hospitals NHS Trust
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Nottingham
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NG5 1PB

Study participating centre

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Headington
Oxford
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OX3 9DU

Study participating centre

Cwm Taf Morgannwg University Health Board
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Coity Rd
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CF31 1RY

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University Hospitals Dorset NHS Foundation Trust
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Bournemouth
United Kingdom
BH7 7DW

Study participating centre

University Hospitals Coventry and Warwickshire NHS Trust

Clifford Bridge Rd

Coventry

United Kingdom

CV2 2DX

Study participating centre

Northumbria Healthcare NHS Foundation Trust

Unit 7-8 Silver Fox Way Cobalt Business Park

Silver Fox Way

Newcastle upon Tyne

United Kingdom

NE27 0QJ

Study participating centre

Leeds Teaching Hospitals NHS Trust

Great George Street

Leeds

United Kingdom

LS1 3EX

Study participating centre

Barnsley Hospital NHS Foundation Trust

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S75 2EP

Study participating centre

Norfolk and Norwich University Hospitals NHS Foundation Trust

Colney Ln

Colney

Norwich

United Kingdom

NR4 7UY

Study participating centre

North Bristol NHS Trust

Southmead Hospital
Southmead Road
Bristol
United Kingdom
BS10 5NB

Study participating centre**Homerton University Hospital NHS Foundation Trust**

Homerton Row
London
United Kingdom
E9 6SR

Study participating centre**Betsi Cadwaladr University Health Board**

Betsi Cadwaladr University Health Board
Ysbyty Gwynedd
Penrhosgarnedd
Bangor
United Kingdom
LL57 2PW

Study participating centre**Gloucestershire Hospitals NHS Foundation Trust**

Great Western Road
Gloucester
United Kingdom
GL1 3NN

Study participating centre**University Hospitals Of Leicester NHS Trust**

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Leicester Royal Infirmary
Infirmary Square
Leicester
United Kingdom
LE1 5WW

Sponsor information

Organisation

University of Bristol

Sponsor details

Research and Enterprise Development
One Cathedral Square
Bristol
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United Kingdom
BS1 5DD
+44 (0)117 42 84021
research-governance@bristol.ac.uk

Sponsor type

University/education

Website

<https://www.bristol.ac.uk/>

ROR

<https://ror.org/0524sp257>

Funder(s)**Funder type**

Government

Funder Name

NIHR Evaluation, Trials and Studies Co-ordinating Centre (NETSCC); Grant Codes: N/K

Results and Publications**Publication and dissemination plan**

Planned publication in a high-impact peer-reviewed journal, conference presentation, publication on website, submission to regulatory authorities

Intention to publish date

28/02/2025

Individual participant data (IPD) sharing plan

The current data sharing plans for this study are unknown and will be 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?
Protocol article		29/10/2021	15/02/2023	Yes	No
HRA research summary			26/07/2023	No	No