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Naltrexone Enhanced Addiction Treatment (NEAT)

Submission date	Recruitment status
08/01/2015	No longer recruiting
Registration date	**Overall study status**
12/01/2015	Completed
Last Edited	**Condition category**
01/02/2019	Mental and Behavioural Disorders

[X] Prospectively registered
[ ] Protocol
[ ] Statistical analysis plan
[X] Results
[ ] Individual participant data

Plain English summary of protocol
Plain English summary under review

Contact information

Type(s)
Scientific

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Additional identifiers

Clinical Trials Information System (CTIS)
2013-002584-25

Protocol serial number
17950

Study information

Scientific Title

Naltrexone Enhanced Addiction Treatment (NEAT): a randomised controlled trial of the clinical and cost-effectiveness of extended-release naltrexone and oral naltrexone

Acronym
NEAT

Study objectives
What is the clinical and cost-effectiveness of enhanced naltrexone (NTX) in the treatment of opioid use disorder? The Naltrexone Enhanced Addiction Therapy for Opioid Use Disorder Trial (NEAT) is the first phase III UK study to coalesce anatagonist medication and behavioural interventions for the treatment of this population.

The study will be implemented in three specialist NHS outpatient addiction clinics in London, Birmingham and Durham (recruitment centres), each with formal links for research trials with a local University.

Three-hundred recently detoxified, formerly dependent heroin users will be randomised to one of three conditions to receive on-site supervised:

1. Thrice-weekly oral active NTX tablets plus placebo extended-release NTX at the start of treatment; or
2. Oral placebo plus active extended-release NTX; or
3. Oral placebo NTX plus placebo extended-release NTX.

Each condition will be delivered over 12 weeks. All participants will receive standard NHS psychological interventions (weekly individual counselling) and a behavioural protocol incentivising clinic attendance to receive trial medication and complete research assessments.

The primary outcome measure is the number of opioid negative urine screening (UDS) tests in treatment (taken thrice weekly during the 12 week treatment phase of the trial; 36 UDS tests in total). In addition to societal focused health-related cost-effectiveness, secondary outcomes include treatment retention/adherence, craving for heroin and cocaine and monitoring of 6-ß-naltrexol (the primary metabolite of NTX). Research worker administered follow-up assessments will be at 16, 24 and 36 weeks after the active 12 week treatment phase.

Ethics approval required
Old ethics approval format

Ethics approval(s)
London - Dulwich Research Ethics Committee, 06/11/2014, ref: 14/LO/1615

Primary study design
Interventional

Study design
Non-randomised; Interventional

Study type(s)
Treatment

Health condition(s) or problem(s) studied
Topic: Mental Health; Subtopic: Addictions; Disease: Addictive Substances– illegal drugs

Interventions
1. Active oral Naltrexone, (O-NTX) and matching placebo, nurse administered (thrice-weekly wk 1-4; Monday and Wednesday: 100mg, Friday: 150mg), patient administered dosing (thrice-weekly wk 5-12; Monday and Wednesday: 100mg, Friday: 150mg)
2. Contingency management element, trial nurse administered, increasing value, non-cash (shop voucher) contingency management element to incentivise attendance to complete study measures (thrice-weekly; wk 1-12)
3. Counselling, 12 sessions of practical, manual-guided, personal goal-setting and relapse-prevention oriented counselling with centre clinician (keyworker, weekly, wk 1-12)
4. Single subcutaneous implant, long-acting form of Naltrexone (herein referred to as XR-NTX, naltrexone, 765mg; iGen/Atral-Cipan) and matching placebo (administered on a day-patient basis by a site doctor at beginning of trial week 1)

Intervention Type
Mixed

Primary outcome(s)
The proportion of opioid negative urines; Timepoint(s): 12 week post randomisation

Key secondary outcome(s)
1. Adherence; Timepoint(s): end of trial
2. Self-reported heroin use via Time-Line Follow-Back (TLFB) interview; Timepoint(s): 16, 24 and 36 weeks
3. Treatment retention; Timepoint(s): end of trial

Completion date
31/05/2016

Eligibility

Key inclusion criteria
Inclusion criteria for the study are intended to be as close to clinical practice as possible. Each participant in the trial must meet all of the following criteria:
1. Is 18 years of age or older
2. Has been given 48 hours to consider and demonstrate verbal understanding of the study patient information material, is able to provide written consent, and can understand and confirm willingness to comply with the protocol.
3. Has a diagnosis of opioid use disorder (heroin; DSM-5: past 12 months), based on the criteria of the Diagnostic and Statistical Manual of Mental Disorders, Fifth Edition, using the MINI Neuropsychiatric Interview (version 6) conducted at baseline.
4. Is completing or has recently completed an inpatient or outpatient treatment for opioid detoxification, or has been completely and continuously abstinent from all opioids for at least seven days.
5. Has no tolerance to opioids, as verified by a negative urine toxicology screening test prior to randomisation (using a Morphine 2000 [opioid class] instant result immunoassay device).
6. Passes a naloxone challenge test (to confirm zero opioid tolerance by demonstrating no clinical sign or subjective report of opioid withdrawal before randomisation and prior to implant

procedure) NB: Individuals failing screening will be allowed to enter screening once more (only) after one month.
7. Is voluntarily seeking opioid antagonist treatment for opioid use disorder.
8. Lives in stable/secure accommodation in the community.
9. Has a personal (mobile/cellular) phone, and is able to nominate at least one locator individual (e.g. a family member, friend or recovery mentor) with a verifiable address and a telephone number to assist with the arrangement of follow-up appointments as required.
10. If female, is not pregnant or breast feeding and agrees to use a birth control method (either oral hormonal contraceptives, barrier [condom or diaphragm], or Nexplanon implant) for the duration of the study

Participant type(s)
Patient

Healthy volunteers allowed
No

Age group
Adult

Lower age limit
18 Years

Sex
All

Key exclusion criteria
Otherwise eligible individuals who meet any of the following criteria will be excluded from the study:
1. Clinically significant medical condition or observed abnormalities on physical examination or laboratory investigation, including but not limited to: Uncontrolled hypertension; Significant heart disease (including angina and myocardial infarction in past 12 months)
Any ECG/cardiovascular abnormality which, in the investigator's judgment, is clinically significant
2. Severe alcohol dependence and/or alcohol intoxication (breathalysed alcohol level <0.35); or alcohol withdrawal
3. Positive naloxone challenge test at randomization (confirming opioid use).
4. Active hepatitis or aspartate aminotransferase, alanine aminotransferase (>3x the upper limit of the normal range)
5. Currently taking oral or depot naltrexone therapy or enrolment in any form of naltrexone therapy within 90 days prior to study screening
6. Current criminal justice involvement with legal proceedings (not including current probationary supervision) and, in the opinion of the clinical worker is expected to fail to complete the study protocol due to re-incarceration or relocation from the centre's catchment area
7. Current (past 30 day) suicidal ideation/plan, or recent (past six months) suicidal ideation or suicide attempt
8. Active, uncontrolled severe mental illness (e.g. psychosis, bipolar I disorder, schizoaffective disorder) and/or a history or evidence of organic brain disease or dementia that would compromise the participant's ability to comply with the study protocol

Date of first enrolment

01/06/2015

Date of final enrolment
31/05/2016

Locations

Countries of recruitment
United Kingdom

England

Study participating centre
King's College London
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Sponsor information

Organisation
King's College London

ROR
https://ror.org/0220mzb33

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

Individual participant data (IPD) sharing plan

IPD sharing plan summary
Not provided at time of registration

Study outputs

Output type	Details	Date created	Date added	Peer reviewed?	Patient-facing?
Results article	results	01/01/2019	01/02/2019	Yes	No
HRA research summary			28/06/2023	No	No