

Biomarkers to target antibiotics and steroid therapy in chronic obstructive pulmonary disease (COPD) exacerbations

Submission date 28/05/2009	Recruitment status No longer recruiting	<input checked="" type="checkbox"/> Prospectively registered <input type="checkbox"/> Protocol
Registration date 30/07/2009	Overall study status Completed	<input type="checkbox"/> Statistical analysis plan <input checked="" type="checkbox"/> Results
Last Edited 12/09/2012	Condition category Respiratory	<input type="checkbox"/> Individual participant data

Plain English summary of protocol
Not provided at time of registration

Contact information

Type(s)
Scientific

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Additional identifiers

Protocol serial number
G0601369

Study information

Scientific Title
The use of biomarkers to direct antibiotic and systemic corticosteroid therapy in chronic obstructive pulmonary disease (COPD) exacerbations: a randomised controlled study

Acronym

BEAT:COPD

Study objectives

Chronic obstructive pulmonary disease (COPD) is a common condition associated with significant morbidity and mortality. It is predicted to be the third leading cause of death worldwide by 2020. COPD exacerbations are an important feature of the disease, accounting for significant morbidity, mortality and health care costs.

COPD exacerbations are associated with bacterial and viral respiratory infections and airway inflammation. Current guidelines advocate the use of oral corticosteroids for patients with a COPD exacerbation who have increased dyspnoea and antibiotics in those with a history of more purulent sputum. A Cochrane review for the use of systemic corticosteroids and antibiotics in COPD exacerbations have shown that corticosteroids increase the rate of recovery following a severe exacerbation, reduce the length of hospital admission and reduce the proportion of patients that have treatment failure. However, it is likely these small corticosteroid-related benefits are confined to a sub-group of patients. Likewise antibiotic therapy in COPD exacerbations is beneficial, with a reduction in short-term mortality and treatment failure; however, the range of response was large and it is estimated that antibiotics are of clinical benefit in only 25 - 50% of COPD exacerbations.

Our inability to identify accurately which patients with a COPD exacerbation should receive antibiotics and or corticosteroids inevitably leading to inappropriate and excessive use of treatment, is the basis of our hypothesis and the use of a single or composite to deliver targeted antibiotic and or corticosteroid therapy at the time of a COPD exacerbation.

As of 20/10/2009 this record was updated after a change to the protocol following MHRA approval. All changes can be found under the relevant section with the above update date. Please note that at this time, the following changes were made:

1. The study design has been updated; the initial study design was: 'Randomised controlled study'
2. The target number of participants has changed; the initial target number of participants was: '136'
3. A placebo arm was added to the interventions section; details of this can be found in the interventions section.

Ethics approval required

Old ethics approval format

Ethics approval(s)

Leicestershire, Northamptonshire and Rutland Research Ethics Committee approved in September 2007 (ref: 07/H0406/157)

Study design

Amended 20/10/2009: A randomised biomarker-driven prednisolone/placebo intervention study

Primary study design

Interventional

Study type(s)

Treatment

Health condition(s) or problem(s) studied

Chronic obstructive pulmonary disease (COPD)

Interventions

Current interventions as of 20/10/2009:

Randomised 12-month parallel group study with two study groups:

1. Standard care therapy group: subjects will receive treatment as decided by a physician and complying with the Global Initiative for Chronic Obstructive Lung Disease (GOLD)/National Institute for Health and Clinical Excellence (NICE) guidelines for management of COPD exacerbations. This may include increasing bronchodilators, a short duration of oral corticosteroids plus or minus antibiotic therapy (according to local hospital microbiological guidelines).
2. Biomarker directed therapy group: subjects will be assigned to 14 days of oral prednisolone or matching placebo as guided by the biomarker and/or 7 days (maximum) of antibiotic therapy.

Randomisation by minimisation: COPD severity, eosinophilic airway inflammation, exacerbation frequency from previous 12 months. Each arm of the study will be followed up for 12 months.

Initial interventions at time of registration:

Randomised 12-month parallel group study with two study groups:

1. Standard care therapy group: subjects will receive treatment as decided by a physician and complying with the Global Initiative for Chronic Obstructive Lung Disease (GOLD)/National Institute for Health and Clinical Excellence (NICE) guidelines for management of COPD exacerbations. This may include increasing bronchodilators, a short duration of oral corticosteroids plus or minus antibiotic therapy (according to local hospital microbiological guidelines).
2. Biomarker directed therapy group: subjects will be assigned to 14 days of oral prednisolone and/or 7 days (maximum) of antibiotic therapy or neither as guided by the biomarker.

Randomisation by minimisation: COPD severity, eosinophilic airway inflammation, exacerbation frequency from previous 12 months. Each arm of the study will be followed up for 12 months.

Intervention Type

Other

Phase

Not Applicable

Primary outcome(s)

1. Proportion of exacerbations treated with antibiotics and corticosteroids
2. Proportion of exacerbations that are associated with a treatment failure
3. Change in health status

Looked at within 3 months of the completion of the study.

Added 20/10/2009:

The study has 80% powering to show equivalence in a minimal change of health status (0.5: measured by the mCRQ).

Key secondary outcome(s)

1. Change in forced expiratory volume in one second (FEV1)
2. Markers of airway inflammation
3. Number of adverse reactions

Looked at within 3 months of the completion of the study.

Completion date

01/09/2010

Eligibility

Key inclusion criteria

1. Provision of informed consent
2. Male or female
3. Aged 40 years or over
4. Diagnosis of COPD
5. Greater than one exacerbation requiring antibiotics and or corticosteroids in the preceding year

Participant type(s)

Patient

Healthy volunteers allowed

No

Age group

Adult

Sex

All

Key exclusion criteria

1. Current active respiratory tuberculosis
2. Upon questioning the patient known human immunodeficiency virus (HIV) infection or positive hepatitis B or C
3. Known inability to produce a sputum sample during the induced sputum procedure
4. Clinically relevant disease or disorder (past or present) which in the opinion of the investigator may either put the subject at risk because of participating in the study or may influence the results of the study or the subject's ability to participate in the study
5. Any clinically relevant lung disease other than COPD
6. Donation of blood within 3 months or during the study (for other than study purpose)
7. Pregnancy or lactation

Date of first enrolment

01/09/2009

Date of final enrolment

01/09/2010

Locations

Countries of recruitment

United Kingdom

England

Study participating centre

Glenfield Hospital

Leicester

United Kingdom

LE3 9QP

Sponsor information**Organisation**

University Hospitals of Leicester NHS Trust (UK)

ROR

<https://ror.org/02fha3693>

Funder(s)**Funder type**

Research council

Funder Name

Medical Research Council (MRC) (UK) (ref: G0601369)

Alternative Name(s)

Medical Research Council (United Kingdom), UK Medical Research Council, Medical Research Committee and Advisory Council, MRC

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	01/07/2012		Yes	No