

Revlimid® Early Stage Poor prognosis Chronic lymphocytic leukaemia (CLL) Trial

Submission date 31/03/2010	Recruitment status No longer recruiting	☒ Prospectively registered ☐ Protocol
Registration date 31/03/2010	Overall study status Completed	☐ Statistical analysis plan ☒ Results
Last Edited 19/05/2022	Condition category Cancer	☐ Individual participant data

Plain English summary of protocol
Not provided at time of registration

Contact information

Type(s)
Scientific

Contact name
Mrs Helen Flight

Contact details
The Christie NHS Foundation Trust
Wilmslow Road
Manchester
United Kingdom
M20 4BX

Additional identifiers

Clinical Trials Information System (CTIS)
2009-011078-14

Protocol serial number
7344

Study information

Scientific Title
A single arm phase II study to investigate the use of Lenalidomide in the treatment of patients with early stage chronic lymphocytic leukaemia (CLL) associated with poor prognostic factors

Acronym

RESPECT

Study objectives

The majority of patients with chronic lymphocytic leukaemia (CLL) are diagnosed with early stage disease (Binet stage A or Rai stage 0/I). Standard management of such patients is observation, and with median age at diagnosis of 72 and median time to progression of greater than 5 - 10 years, many will never require treatment. In contrast, a proportion of patients have more aggressive disease, and over the last decade, a number of molecular factors have been identified that may be used to identify patients with poor prognosis disease. Each is associated with shortened time to treatment (typically less than 3 years in patients with two or more factors), reduced survival, with in the case of p53/ATM inactivation, resistance to treatment.

Whether it is possible to improve the outcome of patients with CLL and adverse prognostic factors by early intervention with treatment is unknown. Several trials in the 1980's demonstrated that treatment of stage A CLL with conventional chemotherapy (chlorambucil) did not alter the natural history of the disease, although none of these studies stratified patients according to risk. The choice of alternative potential therapeutic agents is limited; they should be effective in patients with adverse prognostic factors, have acceptable toxicity, be able to overcome the drug resistance associated with p53/ATM inactivation and ideally be orally administered.

Two recent phase II trials have demonstrated that Lenalidomide is effective in the treatment of relapsed/refractory disease. Importantly, both studies included a high proportion of patients with adverse prognostic factors including p53 inactivation. The principle objective of this study is to investigate the efficacy of Lenalidomide in achieving disease response (complete remission and clearance of minimal residual disease) in patients with poor risk early stage disease, together with assessment of safety and tolerability.

As of 02/05/2012, the anticipated end date of trial has been updated from 01/04/2012 to 24/11/2011.

Ethics approval required

Old ethics approval format

Ethics approval(s)

North West 7 Research Ethics Committee approved on the 27th November 2009 (ref: 09/H1008/122)

Study design

Non-randomised multicentre interventional treatment trial

Primary study design

Interventional

Study type(s)

Treatment

Health condition(s) or problem(s) studied

Topic: National Cancer Research Network; Subtopic: Haematological Oncology; Disease: Leukaemia (chronic)

Interventions

Oral lenalidomide at escalating dose for 3 x 28 day cycles (2.5 mg daily, 5 mg daily, 10 mg daily), then maintenance phase at 10 mg (or maximum tolerated dose).

Intervention Type

Drug

Phase

Phase II

Drug/device/biological/vaccine name(s)

Lenalidomide

Primary outcome(s)

Complete remission with clearance of minimal residual disease (MRD). Response to treatment to be assessed continually, with a more detailed assessment after 6 months of treatment (or earlier if clinically indicated). For patients in complete remission clearance of MRD is assessed every 6 months.

Key secondary outcome(s)

1. Safety and tolerability of treatment, assessed continually throughout treatment by collection of adverse event data, blood results, etc.
2. Event free survival, assessed each time patients are seen - at least once per month during treatment with study drug and then annually once off study drug and in long-term follow-up
3. Time to next treatment, assessed each time patients are seen - at least once per month during treatment with study drug and then annually once off study drug and in long-term follow-up

Completion date

24/11/2011

Eligibility

Key inclusion criteria

1. Binet stage A CLL
2. Two or more risk factors:
 - 2.1. Unmutated IgVH locus (=98% homology to germline sequence)
 - 2.2. CD38 expression (greater than 7%)
 - 2.3. Deletion of chromosome 11q22 (greater than 20% by FISH)
 - 2.4. Deletion of chromosome 17p13 (greater than 10% by FISH)
3. Over 18 years old, either sex
4. Capable to provide written informed consent
5. Eastern Cooperative Oncology Group (ECOG) performance status less than 2
6. Life expectancy greater than 2 years
7. Must agree to not share study lenalidomide with someone else
8. Must agree not to donate blood whilst taking the study drug and for one week after discontinuation of treatment
9. Female subjects of child bearing potential and all male subjects must agree to comply with the stipulations of the pregnancy prevention plan

Participant type(s)

Patient

Healthy volunteers allowed

No

Age group

Adult

Lower age limit

18 years

Sex

All

Key exclusion criteria

1. Current or recent (within the last 1 month) participation in another clinical trial investigation the action of an investigational medicinal product for the treatment of CLL
2. Pregnant or lactating
3. Known positivity for human immunodeficiency virus (HIV) types 1 or 2
4. Prior history of malignancies, other than CLL, unless the subject was treated with curative intent and has been free of the disease for 3 years. Exceptions include the following:
 - 4.1. Basal cell carcinoma of the skin
 - 4.2. Squamous cell carcinoma of the skin
 - 4.3. Carcinoma in situ of the cervix
 - 4.4. Carcinoma in situ of the breast
5. Significantly abnormal renal or hepatic function:
 - 5.1. Creatinine clearance less than 60 ml/min (measured or calculated)
 - 5.2. Serum aspartate aminotransferase (AST) greater than 3 x upper limit of normal (ULN)
 - 5.3. Serum bilirubin greater than 34 µmol/l
6. Laboratory tumour lysis syndrome according to the Cairo-Bishop classification. Subjects may be enrolled when these abnormalities have been corrected.
7. Peripheral neuropathy (grade = 2)
8. Previous treatment for CLL
9. Previous treatment with Thalidomide or immunomodulatory derivative drugs (including lenalidomide)
10. Treatment with corticosteroids (for CLL or other indications) less than 28 days from study entry
11. Evidence of Richter's transformation
12. Unsupported absolute neutrophil count less than $1 \times 10^9/l$ or platelet count less than $50 \times 10^9/l$ not due to CLL
13. Active autoimmune haemolytic anaemia or thrombocytopenia
14. Any other medical or psychological condition that in the view of the investigator would be likely to impact compliance with the protocol or interfere with trial treatment

Date of first enrolment

01/04/2010

Date of final enrolment

24/11/2011

Locations

Countries of recruitment

United Kingdom

England

Study participating centre

The Christie NHS Foundation Trust

550 Wilmslow Road

Manchester

United Kingdom

M20 4BX

Sponsor information

Organisation

Christie NHS Foundation Trust (UK)

ROR

<https://ror.org/03v9efr22>

Funder(s)

Funder type

Industry

Funder Name

Celgene International Sàrl (Switzerland)

Funder Name

Leukaemia Research Fund (LRF) (UK)

Results and Publications

Individual participant data (IPD) sharing plan

Not provided at time of registration

IPD sharing plan summary

Not provided at time of registration

Study outputs

Output type	Details	Date created	Date added	Peer reviewed?	Patient-facing?
Basic results		04/03/2021	19/05/2022	No	No
HRA research summary			28/06/2023	No	No