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A phase I/IIa study combining curcumin (Curcumin C3 Complex, Sabinsa) with standard care FOLFOX chemotherapy in patients with inoperable colorectal cancer

Submission date
28/10/2011

Recruitment status
No longer recruiting

Registration date
28/10/2011

Overall study status
Completed

Last Edited
04/09/2019

Condition category
Cancer

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

Plain English summary of protocol
http://cancerhelp.cancerresearchuk.org/trials/trial-looking-at-curcumin-and-folfox-for-advanced-bowel-cancer

Contact information

Type(s)
Scientific

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Additional identifiers

EudraCT/CTIS number
2011-002289-19

IRAS number

ClinicalTrials.gov number
NCT01490996

Secondary identifying numbers
10672

Study information

Scientific Title
A phase I/IIa study combining curcumin (Curcumin C3 Complex, Sabinsa) with standard care FOLFOX chemotherapy in patients with inoperable colorectal cancer

Acronym
FOLFOX plus curcumin in patients with inoperable colorectal cancer

Study objectives
Phase I/IIa study administering daily oral C3-complex curcumin to patients receiving standard care folinic acid (FOL) fluorouracil (F) and Oxalipatin (OX) (FOLFOX) chemotherapy.

Phase I is a dose-escalation response study to establish a maximum tolerated dose. Phase IIa is a two-armed control study comparing FOLFOX+curcumin with FOLFOX alone. The second phase is randomised. Between 42 and 51 patients will be recruited depending on dose-limiting toxicities observed in phase I. Treatment will last for up to 12 cycles (approx 6 months of chemotherapy). Recruitment is anticipated to take 3 years. Follow-up will include routine CT scans to monitor progression, and ultimately survival times. Median survival approx 18 months and <5% 5 year survival, therefore potential running time 7 years in the event of extremely favourable treatment response.

Ethics approval required
Old ethics approval format

Ethics approval(s)
East Midlands (Derby 1) regional ethics committee , 04/08/2011, ref: 11/EM/0263

Study design
Randomised; Interventional and Observational; Design type: Treatment, Case-controlled study

Primary study design
Interventional

Secondary study design
Randomised controlled trial

Study setting(s)
Hospital

Study type(s)
Treatment

Participant information sheet

http://cancerhelp.cancerresearchuk.org/trials/trial-looking-at-curcumin-and-folfox-for-advanced-bowel-cancer

Health condition(s) or problem(s) studied
Topic: National Cancer Research Network; Subtopic: Colorectal Cancer; Disease: Colon

Interventions
This study is the first to combine daily oral curcumin with standard care FOLFOX-based (5-fluorouracil, folinic acid and oxaliplatin) chemotherapy in colorectal cancer patients with inoperable liver metastases: the CUFOX trial. CUFOX comprises a Phase 1 dose-escalation study (3+3+3 design) to determine an acceptable target dose of curcumin with which to safely proceed to a Phase IIa open-labelled randomised controlled trial. Thirty three participants with histological or cytological confirmation of inoperable colorectal cancer will then be randomised to oxaliplatin-based chemotherapy with the addition of daily oral curcumin at the target dose determined in Phase I, or to standard care oxaliplatin-based chemotherapy alone (recruiting at a ratio of 2:1).

Intervention Type
Drug

Phase
Phase I/II

Drug/device/biological/vaccine name(s)
Folinic acid, fluorouracil, oxaliplatin, curcumin

Primary outcome measure
Safety; Timepoint(s): Real-time adverse event reporting using Common Terminology Criteria for Adverse Events (CTC-AE), diaries and direct consultation.

Secondary outcome measures
1. Biomarker Discovery; Timepoint(s): Samples taken during treatment phase 1-2years. Analysis 3-4 years.
2. Efficacy; Timepoint(s): Measured by response (RECIST) and survival
3. Measurement of systemic curcumin; Timepoint(s): During treatment phase
4. Neurotoxic side-effects; Timepoint(s): 2 weekly neurotoxicity questionnaires
5. Tolerability/Establishing Target Dose; Timepoint(s): Completion dose escalation phase of 3 consecutive patients without dose-limiting toxicitiy 2 cycles

Overall study start date
09/01/2012

Completion date
09/01/2015

Eligibility

Key inclusion criteria
1. Histological or cytological diagnosis of metastatic colorectal cancer
2. Measurable disease by Response Evaluation Criteria in Solid Tumours version 1.1 (RECIST 1.1)

3. Adequate haematological, hepatic and renal function
4. Age more than or equal to 18 years
5. Eastern Cooperative Oncology Group (ECOG) Performance status 0 or 1
6. Patients must have recovered from effects of any recent major surgery
7. Willing to use contraception if applicable
8. Informed consent
9. Life expectancy estimated to be more than 12 weeks; Target Gender: Male & Female ; Lower Age Limit 18 no age limit or unit specified

Participant type(s)
Patient

Age group
Adult

Lower age limit
18 Years

Sex
Both

Target number of participants
Planned Sample Size: 51; UK Sample Size: 51

Total final enrolment
28

Key exclusion criteria
1.Contraindications to FOLFOX chemotherapy: Peripheral neuropathy National Cancer Institute Common Toxicity Criteria (NCI CTC) >1, Liver failure, uncontrolled coronary heart disease, myocardial infarction within the previous 6 months.
2. Unwilling or unable to comply with the study protocol
3. Patients who are pregnant or lactating or contemplating pregnancy. Patients or their partners who become pregnant during the study will be referred to the appropriate experts.
4. Undergone chemotherapy (other than adjuvant for CRC) or participating in another drug study.
5. Previous cancer <5 years (other than colorectal, basal cell carcinoma, in-situ cervical cancer)
6. Major surgery within 4 weeks of starting the study
7. Co-existing active infection or serious concurrent medical condition
8. Significant cardiovascular disease
9. Bone metastases
10. Known brain or leptomeningeal metastases
11. Surgery or hospital admissions for symptomatic intra-abdominal adhesions
12. Active endoscopically proven peptic ulcer disease or colitis

Date of first enrolment
09/01/2012

Date of final enrolment
09/01/2015

Locations

Countries of recruitment
England

United Kingdom

Study participating centre
Leicester Royal Infirmary
Leicester
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LE1 5WW

Sponsor information

Organisation
University of Leicester (UK)

Sponsor details
Department of Cancer Studies and Molecular Medicine
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Sponsor type
University/education

ROR
https://ror.org/04h699437

Funder(s)

Funder type
Charity

Funder Name
Bowel Disease Research Foundation of ACPGBI (UK)

Funder Name
Cancer Research UK (CRUK) (UK)

Alternative Name(s)
CR_UK, Cancer Research UK - London, CRUK

Funding Body Type
Private sector organisation

Funding Body Subtype
Other non-profit organizations

Location
United Kingdom

Funder Name
Hope Against Cancer (UK)

Funder Name
Royal College of Surgeons of England (UK)

Alternative Name(s)
RCS

Funding Body Type
Private sector organisation

Funding Body Subtype
Associations and societies (private and public)

Location
United Kingdom

Results and Publications

Publication and dissemination plan
Not provided at time of registration

Intention to publish date

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?
Protocol article	protocol	24/03/2015		Yes	No
Results article	results	01/07/2019	04/09/2019	Yes	No
HRA research summary			28/06/2023	No	No