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A Mechanistic Study Of Mifamurtide (MTPPE) In Patients With Metastatic And/Or Recurrent Osteosarcoma

Submission date 18/09/2014	**Recruitment status** No longer recruiting	[_] Prospectively registered [_] Protocol [_] Statistical analysis plan [X] Results [_] Individual participant data
Registration date 18/09/2014	**Overall study status** Completed	
Last Edited 11/06/2019	**Condition category** Cancer	

Plain English summary of protocol
http://www.cancerresearchuk.org/about-cancer/trials/a-trial-of-mifamurtide-for-advanced-osteosarcoma-memos

Contact information

Type(s)
Scientific

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Additional identifiers

Clinical Trials Information System (CTIS)
2012-000615-84

ClinicalTrials.gov (NCT)
NCT02441309

Protocol serial number
16801

Study information

Scientific Title
A Mechanistic Study Of Mifamurtide (MTPPE) In Patients With Metastatic And/Or Recurrent Osteosarcoma

Acronym
MEMOS: a Eurosarc Study of Mifamurtide in advanced osteosarcoma

Study objectives
This is a Bayesian designed multi-arm, multi-centre open-label phase II study in patients with metastatic and/or recurrent osteosarcoma, which will investigate why some patients with osteosarcoma may respond better than others to mifamurtide given alone or in combination with ifosfamide.

Ethics approval required
Old ethics approval format

Ethics approval(s)
South Central - Oxford C Research Ethics Committee, 13/06/2014, ref: 14/SC/0255

Study design
Both; Interventional; Design type: Treatment

Primary study design
Interventional

Study type(s)
Treatment

Health condition(s) or problem(s) studied
Topic: Cancer; Subtopic: Sarcoma; Disease: Bone

Interventions
Patients with relapsed or metastatic osteosarcoma will be divided into three treatment groups (Arms). Depending on their current disease status, patients may be either Registered to Arm A (resectable group), to receive Mifamurtide alone; or Randomised to Arm B/C (non-resectable group), to receive mifamurtide in combination with ifosfamide.
Arm A - Mifamurtide alone; Arm B - Ifosfamide alone for 6 weeks then Ifosfamide + mifamurtide for 6 weeks, then mifamurtide alone for 30 weeks; Arm C - Ifosfamide + mifamurtide for 12 weeks then mifamurtide alone for 24 weeks. All participants will receive 36 weeks or more of mifamurtide.
Biopsies (or resected tumour samples) will be obtained before and after 6 weeks of therapy interval in order to determine the pharmacodynamic endpoints. The target sample size is 40 patients. An interim analysis will be performed for the primary efficacy endpoint.

Intervention Type

Drug

Phase
Phase II

Drug/device/biological/vaccine name(s)
Mifamurtide, Ifosfamide

Primary outcome(s)
Objective Radiological response; Timepoint(s): Pre-treatment, after 6, 12, 18, 24 & 36 weeks of treatment

Key secondary outcome(s))
1. Biological response; Timepoint(s): Pre-treatment and after 6 weeks of treatment
2. Disease specific overall survival; Timepoint(s): End of trial
3. Progression free survival on serial CT scan; Timepoint(s): Pre-treatment, after 6, 12, 18, 24 and 36 of weeks of treatment
4. Safety and tolerability on CTCAE Criteria (v4.0); Timepoint(s): Throughout trial treatment

Completion date
11/03/2016

Eligibility

Key inclusion criteria
1. Relapsed osteosarcoma (first, second, third or any relapse, patient has recovered from chemotherapy and any other investigational drug/agent treatment, radiotherapy or surgical procedure).
2. Histological confirmed diagnosis of osteosarcoma at original presentation.
3. Tumour at biopsy accessible or resectable site.
4. Progressive disease documented by imaging within 3 months of entry into the trial.
5. At least one measurable lesion on CT scan (RECIST) performed in past 21 days prior to trial entry.
6. Male or female, age = 16 years to 65 (or =18 based on institutional practice for Teenage and Young Adult Cancer patients).
7. Life expectancy of at least 3 months.
8. WHO performance score of 0 - 2.
9. The patient is willing and able to comply with the protocol and scheduled follow-up visits and examinations.
10. Written (signed and dated) informed consent.
11. Cardiac shortening fraction = 28% or ejection fraction = 45%
12. Renal function is adequate for ifosfamide treatment (GFR as per table below, other renal function screening tests as per local practice)
13. Haematological and biochemical indices within the ranges detailed in the protocol

Participant type(s)
Patient

Healthy volunteers allowed
No

Age group
Adult

Lower age limit
18 years

Sex
All

Key exclusion criteria
1. Pregnant or breastfeeding woman. Men or women of childbearing potential unless effective methods of contraception are used during study treatment and for at least 7 days after the last mifamurtide dose.
2. Previous treatment with mifamurtide or a mifamurtide like drug* in a clinical trial setting for the treatment of metastatic and/or recurrent osteosarcoma in the six months prior to registration.
3. Contraindications to lung biopsies
4. Hypersensitivity to ifosfamide or any component of the formulation.
5. Previously diagnosed brain metastases.
6. Significant active cardiac disease including: uncontrolled high blood pressure (no greater than 2 standard deviations above the mean for age for systolic blood pressure (SBP) and diastolic blood pressure (DBP), unstable angina, congestive heart failure, myocardial infarction within the previous 6 months, or serious cardiac arrhythmias and with a history of pericarditis and myocarditis
7. Treatment with any other investigational agent, or participation in another interventional clinical trial within 21
days prior to enrolment.
8. Major surgery within 21 days prior first study biopsy
9. Currently taking of high-dose nonsteroidal antiinflammatory drugs (NSAIDs) or corticosteroid treatment
10. Concurrent use of ciclosporin or other calcineurin inhibitors.
11. Any psychological, social or medical condition, physical examination finding or a laboratory abnormality that the Investigator considers would make the patient a poor trial candidate or could interfere with protocol compliance or the interpretation of trial results.
12. Any other active malignancy, with the exception of adequately treated conebiopsied in situ carcinoma of the
cervix uteri and non-melanoma skin lesions.
13. Patients who are known to be serologically positive for Hepatitis B, Hepatitis C or HIV.
* mifamurtide-like drugs include GMCSF, interferon and other macrophage activating molecules.

Date of first enrolment
12/09/2014

Date of final enrolment
11/03/2016

Locations

Countries of recruitment
United Kingdom

England

Study participating centre
Oncology Clinical Trials Office (OCTO)
Oxford
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Sponsor information

Organisation
University of Oxford (UK)

ROR
https://ror.org/052gg0110

Funder(s)

Funder type
Government

Funder Name
European Commission

Alternative Name(s)
European Union, Comisión Europea, Europäische Kommission, EU-Kommissionen, Euroopa Komisjoni, EC, EU

Funding Body Type
Government organisation

Funding Body Subtype
National government

Location

Funder Name
Millenium Pharmaceuticals Inc. (USA)

Results and Publications

Individual participant data (IPD) sharing plan

The data sharing plans for the current study are unknown and will be made 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?
Basic results				No	No
HRA research summary			28/06/2023	No	No
Participant information sheet	Participant information sheet	11/11/2025	11/11/2025	No	Yes