

A phase I study to evaluate the safety and pharmacokinetics of ONX 0801 in advanced solid tumours

Submission date 12/06/2009	Recruitment status No longer recruiting	☒ Prospectively registered ☐ Protocol
Registration date 30/06/2009	Overall study status Completed	☐ Statistical analysis plan ☐ Results
Last Edited 12/12/2017	Condition category Cancer	☐ Individual participant data ☐ Record updated in last year

Plain English summary of protocol

Not provided at time of registration

Contact information

Type(s)

Scientific

Contact name

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Contact details

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Additional identifiers

Protocol serial number

2009-001

Study information

Scientific Title

A phase I, open-label, dose-finding study to evaluate the safety and pharmacokinetics of ONX 0801, a novel alpha-folate receptor-mediated thymidylate synthase inhibitor, in patients with advanced solid tumours

Study objectives

Is ONX 0801 tolerable and safe in cancer patients and can a dose be identified which inhibits tumour cell growth in future clinical studies?

Ethics approval required

Old ethics approval format

Ethics approval(s)

Royal Marsden Hospital Ethics Committee and the Hammersmith Ethics Committee – submission pending, planned for June 2009

Primary study design

Interventional

Study design

Phase I open-label dose-finding study

Study type(s)

Treatment

Health condition(s) or problem(s) studied

Advanced solid tumours

Interventions

Cohorts of 3 to 6 patients will receive ONX 0801 at escalating doses until a maximum tolerated dose (MTD) is determined. Each patient will receive a 3-hour intravenous (IV) infusion of ONX 0801 weekly (i.e., on days 1, 8, and 15) of repeated 21-day treatment cycles.

Contact details for patient information material:

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Intervention Type

Drug

Phase

Phase I

Drug/device/biological/vaccine name(s)

ONX 0801

Primary outcome(s)

1. To determine the maximum tolerated dose (MTD) and/or recommended phase II dose (RP2D) of ONX 0801 based on dose limiting toxicities (DLTs) occurring within cycle 1
2. To characterise the safety profile of ONX 0801

Key secondary outcome(s)

1. Pharmacokinetics (PK) of ONX 0801:

Blood samples will be collected according to the following schedule:

1.1. Cycle 1, Day 1: predose and 30 minutes and 1, 2, 3, 3.5, 4, 6, 8, 12, 24 (Day 2), 48 (Day 3), and 72 (Day 4) hours following the start of the infusion

1.2. Cycle 1, Days 8 and 15: predose and 3 hours following the start of the infusion

1.3. Cycle 2, Days 1 and 8: predose and 3 hours following the start of the infusion

2. Pharmacodynamics of ONX 0801:

2.1. Blood samples will be collected according to the following schedule: predose and 4, 8, 24 (Day 2), 48 (Day 3), and 72 (Day 4) hours following the start of infusion in Cycle 1, Day 1 and approximately every 6 - 9 weeks during the course of the study

2.2. Tissue samples may be collected predose and up to 72 hours following the start of the infusion in Cycle 1, Day 1

2.3. 18FLT-PET scans may be performed predose and between 16 to 48 hours following the start of the infusion in Cycle 1, Day 1

3. Identifying a biologically effective dose (BED) equal to or lower than the MTD and/or RP2D of ONX 0801

4. Assess the preliminary antitumour activity of ONX 0801

Completion date

30/03/2011

Eligibility

Key inclusion criteria

1. Histologically or cytologically proven solid tumours, including lymphomas. Patients must have disease which has failed standard therapy or for which no standard curative therapy exists.

2. Greater than or equal to 18 years of age, either sex

3. Eastern Cooperative Oncology Group performance status (ECOG PS) less than or equal to 2

4. Life expectancy greater than or equal to 12 weeks

5. Measurable (as defined by Response Evaluation Criteria in Solid Tumours [RECIST version 1.1]) or evaluable (based on radiological assessments or tumour markers) disease

6. Recovered (i.e., to National Cancer Institute [NCI] Common Terminology Criteria for Adverse Events [CTCAE] Version 3.0 Grade less than or equal to 1) from all toxicities associated with previous chemotherapy or radiotherapy (exception: patients may enter with continuing alopecia irrespective of CTCAE grade). The following intervals between starting last treatment and starting ONX 0801 must elapse:

6.1. Chemotherapy (see exception below): at least 4 weeks

6.2. Mitomycin C or a nitrosourea: at least 6 weeks

6.3. Targeted therapy: at least 2 weeks or 2 half-lives, whichever is longer

6.4. Biologics: at least 4 weeks

6.5. Radiotherapy: at least 4 weeks

7. Normal organ function

8. Normal electrocardiogram (ECG)

9. Archival tumour tissue available

Participant type(s)

Patient

Healthy volunteers allowed

No

Age group

Adult

Lower age limit

18 Years

Sex

All

Key exclusion criteria

1. Pregnant women, women who are lactating, or women of childbearing potential who are not currently on effective means of birth control
2. History of QT/QTc prolongation, clinically significant ventricular tachycardia, ventricular fibrillation, heart block, myocardial infarction within 1 year, congestive heart failure New York Heart Association Class III or IV, unstable angina, angina within 6 months, or other evidence of clinically significant coronary artery disease
3. Active, ongoing infection, including viral hepatitis
4. Undergone major surgery within the last 4 weeks
5. Organ transplant recipients
6. New brain metastasis. Patients with treated (surgically excised or irradiated) and stable brain metastases are eligible as long as the treatment was at least 4 weeks prior to initiation of study drug and baseline brain computed tomography (CT) with contrast or magnetic resonance imaging (MRI) within 2 weeks of initiation of study drug is negative for new brain metastases.
7. Patients who have been on other experimental clinical trials of investigational agents within the last 28 days

Date of first enrolment

30/09/2009

Date of final enrolment

30/03/2011

Locations**Countries of recruitment**

United Kingdom

United States of America

Study participating centre

2100 Powell Street

Emeryville

United States of America

94608

Sponsor information

Organisation

Onyx Pharmaceuticals (USA)

ROR

<https://ror.org/03g03ge92>

Funder(s)

Funder type

Industry

Funder Name

Onyx Pharmaceuticals (USA)

Alternative Name(s)

Funding Body Type

Private sector organisation

Funding Body Subtype

For-profit companies (industry)

Location

United States of America

Results and Publications

Individual participant data (IPD) sharing plan

IPD sharing plan summary

Not provided at time of registration