

Comparing a combination gemcitabine and Vandetanib therapy with gemcitabine therapy alone in locally advanced or metastatic Pancreatic carcinoma

Submission date 10/08/2011	Recruitment status No longer recruiting	<input checked="" type="checkbox"/> Prospectively registered <input type="checkbox"/> Protocol
Registration date 10/08/2011	Overall study status Completed	<input type="checkbox"/> Statistical analysis plan <input checked="" type="checkbox"/> Results
Last Edited 06/03/2017	Condition category Cancer	<input type="checkbox"/> Individual participant data

Plain English summary of protocol

<http://cancerhelp.cancerresearchuk.org/trials/a-trial-comparing-gemcitabine-vandetanib-gemcitabine-alone-pancreatic-cancer-vip>

Study website

http://www.lctu.org.uk/trial/trial_info.asp?id=72&tgcode=4&menuid=30

Contact information

Type(s)

Scientific

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Additional identifiers

EudraCT/CTIS number

2010-021951-26

IRAS number

ClinicalTrials.gov number

Secondary identifying numbers

9908

Study information

Scientific Title

A prospective, phase II, double blinded, multicentre, randomised clinical trial comparing combination gemcitabine and vandetanib therapy with gemcitabine therapy alone in locally advanced or metastatic pancreatic carcinoma

Acronym

ViP

Study objectives

To assess whether survival times for patients receiving gemcitabine plus vandetanib are longer than for those patients receiving gemcitabine alone as first line treatment for advanced pancreatic cancer.

Ethics approval required

Old ethics approval format

Ethics approval(s)

First MREC, 04/07/2011, ref: 11/LO/0097

Study design

Prospective phase II placebo-controlled multicentre randomised clinical trial

Primary study design

Interventional

Secondary study design

Randomised controlled trial

Study setting(s)

GP practice

Study type(s)

Treatment

Participant information sheet

Not available in web format, please use the contact details to request a patient information sheet

Health condition(s) or problem(s) studied

Upper gastro-intestinal cancer, pancreatic cancer

Interventions

The planned treatment duration per patient will be until progression of disease, unacceptable toxicity or withdrawal of consent. The end of the study will be 12 months after the recruitment of the last patient. Patients who stop treatment before having developed progressive disease (PD) will be assessed every 6 weeks for response until PD occurs. Subjects will be randomised equally to two arms:

1. Arm A (standard therapeutic arm): Placebo orally once a day continuously together with Gemcitabine 1000mg/m² 2 weekly as a 30 minute infusion for 7 consecutive weeks, followed by a one week break, followed by gemcitabine 1000mg/m² 2 weekly as a 30 minute infusion for 3 weeks followed by a one week break in subsequent cycles.
2. Arm B: vandetanib orally once a day continuously at 300 mg/day together with Gemcitabine 1000mg/m² 2 weekly as a 30 minute infusion for 7 consecutive weeks, followed by a one week break, followed by Gemcitabine 1000mg/m² 2 weekly as a 30 minute infusion for 3 weeks followed by a one week break in subsequent cycles.

Gemcitabine administration: Gemcitabine 1000mg/m² 2 weekly as a 30 minute infusion for 7 consecutive weeks, followed by a one week break, followed by Gemcitabine 1000mg/m² 2 weekly as a 30 minute infusion for 3 weeks followed by a one week break in subsequent cycles. Vandetanib administration, 300mg/day orally continuously in the form of a white tablet. Followed up at 24 months

Intervention Type

Drug

Phase

Phase II

Drug/device/biological/vaccine name(s)

Gemcitabine, vandetanib

Primary outcome measure

1. Assessment of survival times between the two arms
2. Assess survival between subjects on arm A compared to arm B

Secondary outcome measures

1. Progression-free survival time (PFS)
2. Objective response rate
3. Disease control rate
4. Toxicity and safety

Overall study start date

01/09/2011

Completion date

01/09/2012

Eligibility

Key inclusion criteria

1. Age > 18 years
2. Histologically or cytologically proven pancreatic ductal adenocarcinoma or undifferentiated

carcinoma of the pancreas

3. Locally advanced or metastatic disease precluding curative surgical resection or definitive locally directed therapies such as chemo radiation. Patients who have relapsed following previously resected pancreatic cancer can be included.

4. Contrast enhanced computerised tomography (CT) scan of the thorax, abdomen and pelvis within 28 days prior to commencing treatment

5. Unidimensionally measurable disease as shown by CT scan, in accordance with the Response Evaluation Criteria In Solid Tumours (RECIST) guidelines (version 1.1).

6. ECOG performance status 0, 1 or 2 where the investigator feels that treatment with combination chemotherapy, for example FOLFIRINOX, is not appropriate

7. Platelets $>100 \times 10^9/l$; WBC $> 3 \times 10^9/l$; neutrophils $> 1.5 \times 10^9/l$ at entry

8. Documented Life expectancy > 3 months

9. Informed written consent

10. Male and female participants

Participant type(s)

Patient

Age group

Adult

Lower age limit

18 Years

Sex

Both

Target number of participants

Planned Sample Size: 120; UK Sample Size: 120

Key exclusion criteria

1. Laboratory results: Serum bilirubin $>1.5 \times$ the upper limit of reference range (ULRR). Creatinine clearance < 30 mL/minute (calculated by Cockcroft-Gault formula).

Potassium, <4.0 mmol/L despite supplementation; or above the CTCAE grade 1 upper limit.

Magnesium below the normal range despite supplementation, or above the CTCAE grade 1 upper limit. Serum corrected calcium above the CTCAE grade 1 upper limit. In cases where the serum calcium is below the normal range despite supplementation.

Alanine aminotransferase (ALT) or aspartate aminotransferase (AST) > 2.5 ULRR or alkaline phosphatase (ALP) $>2.5 \times$ ULRR, or $> 5 \times$ ULRR if judged by the investigator to be related to liver metastases.

2. Medical or psychiatric conditions compromising informed consent

3. Intracerebral metastases or meningeal carcinomatosis

4. Major surgery within 4 weeks or incompletely healed surgical incision before starting study therapy

5. Evidence of severe or uncontrolled systemic disease or any concurrent condition which in the Investigators opinion makes it undesirable for the patient to participate in the trial or which would jeopardize compliance with the protocol

6. Clinically significant cardiovascular event (e.g. myocardial infarction, superior vena cava syndrome (SVC), New York Heart Association (NYHA) classification of heart disease ≥ 2 within 3 months before entry; or presence of cardiac disease that, in the opinion of the Investigator, increases the risk of ventricular arrhythmia

7. History of arrhythmia (multifocal premature ventricular contractions [PVCs], bigeminy,

- trigeminy, ventricular tachycardia, or uncontrolled atrial fibrillation), which is symptomatic or requires treatment (CTCAE grade 3) or asymptomatic sustained ventricular tachycardia. Atrial fibrillation, controlled on medication is not excluded
8. QTc prolongation with other medications that required discontinuation of that medication
 9. Congenital long QT syndrome or 1st degree relative with unexplained sudden death under 40 years of age
 10. Presence of left bundle branch block (LBBB)
 11. QTc with Bazetts correction that is un-measurable, or 480 msec on screening ECG
(Note: If a subject has a QTc interval 480 msec on screening ECG, the screen ECG may be repeated twice [at least 24 hours apart]. The average QTc from the three screening ECGs must be <480 msec in order for the subject to be eligible for the study.) Patients who are receiving a drug that has a risk of inducing Torsades-de-Pointes are excluded if QTc is = 460 msec.
 12. Any concurrent medication with a known risk of inducing Torsades-de-Pointes, that in the investigators opinion cannot be discontinued, are allowed; however, these patients must be monitored closely (please see section 4.2).
 13. Concomitant medications that are potent inducers (rifampicin, rifabutin, phenytoin, carbamazepine, phenobarbital and St. John's Wort) of CYP3A4 function.
 14. Hypertension not controlled by medical therapy (systolic blood pressure greater than 160 mm Hg or diastolic blood pressure greater than 100 mm Hg).
 15. Currently active diarrhoea that may affect the ability of the patient to absorb the vandetanib or tolerate diarrhoea secondary to vandetanib should that occur as a side effect.
 16. Malabsorption syndrome which may impair the absorption of vandetanib (partial gastrectomy, small bowel resection), This may include previous partial gastrectomy and small bowel resection or active Crohns disease, ulcerative colitis.
 17. Pregnancy or breast feeding.
 18. Previous chemotherapy for locally advanced and metastatic disease. Adjuvant chemotherapy for resected pancreatic cancer will be permitted provided that chemotherapy was completed > 12 months previously.
 19. Radiotherapy within the last 4 weeks prior to start of study treatment.
 20. Concurrent malignancies or invasive cancers diagnosed within past 5 years except for adequately treated basal cell carcinoma of the skin, in situ carcinoma of the uterine cervix or resected pancreatic cancer.
 21. Chemotherapy directed at tumour apart from that described in this protocol.
 22. All men or women of reproductive potential, unless using at least two contraceptive precautions, one of which must be a condom..

Date of first enrolment

01/09/2011

Date of final enrolment

01/09/2012

Locations

Countries of recruitment

England

United Kingdom

Study participating centre
University of Liverpool
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Sponsor information

Organisation
University of Liverpool (UK)

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Research & Development
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Sponsor type
University/education

Website
<http://www.liv.ac.uk/>

ROR
<https://ror.org/04xs57h96>

Funder(s)

Funder type
Industry

Funder Name
AstraZeneca (UK)

Alternative Name(s)
AstraZeneca PLC, Pearl Therapeutics

Funding Body Type
Government organisation

Funding Body Subtype
For-profit companies (industry)

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?
Results article	results	01/04/2017		Yes	No
HRA research summary			28/06/2023	No	No