

# A Phase II study of axitinib in patients with metastatic renal cell cancer unsuitable for nephrectomy

|                                        |                                                   |                                                                                                              |
|----------------------------------------|---------------------------------------------------|--------------------------------------------------------------------------------------------------------------|
| <b>Submission date</b><br>26/09/2012   | <b>Recruitment status</b><br>No longer recruiting | <input checked="" type="checkbox"/> Prospectively registered<br><input type="checkbox"/> Protocol            |
| <b>Registration date</b><br>26/09/2012 | <b>Overall study status</b><br>Completed          | <input type="checkbox"/> Statistical analysis plan<br><input type="checkbox"/> Results                       |
| <b>Last Edited</b><br>18/12/2024       | <b>Condition category</b><br>Cancer               | <input type="checkbox"/> Individual participant data<br><input type="checkbox"/> Record updated in last year |

## Plain English summary of protocol

<https://www.cancerresearchuk.org/about-cancer/find-a-clinical-trial/a-trial-looking-axitinib-kidney-cancer-spread-a-predict>

## Contact information

### Type(s)

Scientific

### Contact name

Ms Rebecca Lewis

### Contact details

Institute of Cancer Research Clinical Trials & Statistics Unit (ICR-CTSU)

Section of Clinical Trials

Sir Richard Doll Building

15 Cotswold Road

Sutton

United Kingdom

SM2 5NG

-

apredict-icrctsu@icr.ac.uk

## Additional identifiers

### ClinicalTrials.gov (NCT)

NCT01693822

### Clinical Trials Information System (CTIS)

2011-004562-16

## **Integrated Research Application System (IRAS)**

98117

### **Protocol serial number**

12404

## **Study information**

### **Scientific Title**

A Phase II study of axitinib in patients with metastatic renal cell cancer unsuitable for nephrectomy

### **Acronym**

A-PREDICT

### **Study objectives**

A-PREDICT is a single arm, single agent, open label, multicentre, phase II study of axitinib in patients with metastatic renal cell carcinoma of predominant clear cell histology and unsuitable for debulking nephrectomy (as judged by the treating clinician).

Patients who have provided consent and have satisfied the eligibility criteria will be registered into the trial. The starting dose of axitinib will be 5 mg twice daily by mouth, escalating to a maximum of 10mg twice daily by mouth according to tolerability of treatment, for as long as patients are deriving clinical benefit. Treatment will be paused for one week prior to the week 9 biopsy. Disease progression will be evaluated according to RECIST v1.1 criteria 8 weeks after commencing treatment, at 8 weekly intervals to 6 months and 3 monthly thereafter. Blood and tumour tissue samples will be taken prior to and during therapy to evaluate biomarkers of treatment response. Nephrectomy will be carried out on any patient who becomes suitable in the opinion of the treating clinician during the course of the trial. Where possible, tissue samples will be taken from resected specimens. Response to axitinib in marker lesions will be correlated with changes in biomarkers.

### **Ethics approval required**

Old ethics approval format

### **Ethics approval(s)**

First MREC, 08/05/2012 ref: 12/LO/0639

### **Primary study design**

Interventional

### **Study design**

Non-randomised interventional trial

### **Study type(s)**

Treatment

### **Health condition(s) or problem(s) studied**

Renal cancer

## Interventions

Oral axitinib twice a day. 5mg starting dose escalated to maximum of 10mg until disease progression

## Intervention Type

Drug

## Phase

Phase II

## Drug/device/biological/vaccine name(s)

Axitinib

## Primary outcome(s)

Freedom from progression at 6 months, measured at 6 months after start of treatment

## Key secondary outcome(s)

No secondary outcome measures

## Completion date

28/02/2022

## Eligibility

### Key inclusion criteria

1. Histologically confirmed metastatic renal cell carcinoma of predominant clear cell histology
2. Unsuitable for nephrectomy as judged by treating clinician(s)
3. Not suitable for watch and wait policy as determined by treating clinician(s)
4. No prior systemic therapy for renal cell carcinoma
5. Measurable metastatic disease using RECIST v1.1
6. 18 years of age or older
7. Life expectancy of 12 weeks or greater
8. ECOG performance status 0 or 1
9. Adequate organ function as defined by serum aspartate transaminase (AST) and serum alanine transaminase (ALT)  $\leq 2.5 \times$  upper limit of normal (ULN), or AST and ALT  $\leq 5 \times$  ULN if liver function abnormalities are due to liver metastases; total serum bilirubin  $\leq 1.5 \times$  ULN
10. Adequate haematological function as defined by absolute neutrophil count (ANC)  $\geq 1500/\mu\text{L}$ , platelets  $\geq 75,000/\mu\text{L}$ , haemoglobin  $\geq 9.0 \text{ g/dL}$  and prothrombin time (PT)  $\leq 1.5 \times$  ULN
11. Serum creatinine  $\leq 1.5 \times$  ULN or calculated creatinine clearance  $\geq 60 \text{ mL/min}$ ;
12. Urinary protein  $< 2+$  by urine dipstick. If dipstick is  $\geq 2+$  then a 24-hour urine collection can be done and the patient may enter only if urinary protein is  $< 2\text{g}$  per 24 hours.
13. No evidence of pre-existing uncontrolled hypertension as documented by 2 baseline blood pressure readings taken at least 1 hour apart. The baseline systolic blood pressure readings must be  $\leq 140 \text{ mm Hg}$ , and the baseline diastolic blood pressure readings must be  $\leq 90 \text{ mm Hg}$ . Patients whose hypertension is controlled by antihypertensive therapies are eligible.
14. Women of childbearing potential must have a negative serum or urine pregnancy test within 3 days prior to treatment.
15. Willingness and ability to comply with scheduled visits, treatment plans, laboratory tests, and

other study procedures, including tumour biopsies.

16. Written informed consent

17 Male or female participants

**Participant type(s)**

Patient

**Healthy volunteers allowed**

No

**Age group**

Adult

**Lower age limit**

18 Years

**Sex**

All

**Total final enrolment**

65

**Key exclusion criteria**

1. The presence of intracranial disease, unless there has been radiological evidence of stable intracranial disease >6 months. In the case of a solitary brain metastasis which has been resected, there must be evidence of a disease-free interval of at least 3 months postsurgery. All patients previously treated for brain metastases must be stable off corticosteroid therapy for at least 28 days.

2. The presence of active second malignancy. Patients will be eligible if they have adequately treated basal cell carcinoma, squamous cell skin cancer, in situ cervical cancer, stable prostate cancer or if treated with curative intent for any other cancer with no evidence of disease for 2 years.

3. Women who are pregnant or are breastfeeding. Female patients must be surgically sterile, be postmenopausal, or must agree to use effective contraception during the period of therapy. All female patients with reproductive potential must have a negative pregnancy test (serum or urine) prior to enrolment.

4. Male patients must be surgically sterile or must agree to use effective contraception during the period of therapy.

5. Current signs or symptoms of severe progressive or uncontrolled hepatic, endocrine, pulmonary disease other than directly related to RCC.

6. Gastrointestinal abnormalities including:

6.1. Inability to take oral medication

6.2. Requirement for intravenous alimentation

6.3. Prior surgical procedures affecting absorption including total gastric resection

6.4. Treatment for active peptic ulcer disease in the past 6 months

6.5. Active gastrointestinal bleeding, unrelated to cancer, as evidenced by hematemesis, hematochezia or melena in the past 3 months without evidence of resolution documented by endoscopy or colonoscopy;

6.6. Malabsorption syndromes.

7. Current use or anticipated need for treatment with drugs that are known potent CYP3A4 inhibitors

8. Current use or anticipated need for treatment with drugs that are known CYP3A4 or CYP1A2 inducers
9. Requirement of anticoagulant therapy with oral vitamin K antagonists. Low-dose anticoagulants for maintenance of patency of central venous access device or prevention of deep venous thrombosis is allowed. Therapeutic use of low molecular weight heparin is allowed.
10. Active seizure disorder, spinal cord compression, or carcinomatous meningitis.
11. Any of the following within 12 months prior to study entry: myocardial infarction, uncontrolled angina, coronary/peripheral artery bypass graft, symptomatic congestive heart failure, cerebrovascular accident or transient ischemic attack.
12. Deep vein thrombosis or pulmonary embolism within 6 months prior to study entry.
13. Known human immunodeficiency virus (HIV) or acquired immunodeficiency syndrome (AIDS)-related illness.

**Date of first enrolment**

28/09/2012

**Date of final enrolment**

27/09/2014

## Locations

**Countries of recruitment**

United Kingdom

England

**Study participating centre**

Institute of Cancer Research Clinical Trials & Statistics Unit (ICR-CTSU)

Sutton

United Kingdom

SM2 5NG

## Sponsor information

**Organisation**

Institute of Cancer Research (UK)

**ROR**

<https://ror.org/043jzw605>

## Funder(s)

**Funder type**

Industry

**Funder Name**

Pfizer UK

**Alternative Name(s)**

Pfizer Ltd, Pfizer Limited

**Funding Body Type**

Private sector organisation

**Funding Body Subtype**

For-profit companies (industry)

**Location**

United Kingdom

## Results and Publications

**Individual participant data (IPD) sharing plan**

The datasets generated during and/or analysed during the current study are/will be available upon request from Rebecca Lewis (apredict-icrctsu@icr.ac.uk). Clinical data are available for sharing subject to completion of a data sharing application form, approval by the trial oversight committees and completion of a data sharing agreement. As part of the review the trialists would consider whether the existing trial consent covers the application, what anonymisation will be required and whether separate ethics approval would be required.

**IPD sharing plan summary**

Available on request

**Study outputs**

| Output type                           | Details       | Date created | Date added | Peer reviewed? | Patient-facing? |
|---------------------------------------|---------------|--------------|------------|----------------|-----------------|
| <a href="#">HRA research summary</a>  | Study website | 11/11/2025   | 28/06/2023 | No             | No              |
| <a href="#">Plain English results</a> |               |              | 26/10/2022 | No             | Yes             |
| <a href="#">Study website</a>         |               |              | 11/11/2025 | No             | Yes             |