A Study to Evaluate Pazopanib as an Adjuvant Treatment for Localized Renal Cell Carcinoma (RCC) (PROTECT)

This study is currently recruiting participants.

Verified July 2012 by GlaxoSmithKline

First Received on October 14, 2010. Last Updated on July 12, 2012

Purpose

This randomized Phase III study is to evaluate whether pazopanib compared with placebo can prevent or delay recurrence of kidney cancer in patients with moderately high or high risk of developing recurrence after undergoing kidney cancer surgery.

<table>
<thead>
<tr>
<th>Condition</th>
<th>Intervention</th>
<th>Phase</th>
</tr>
</thead>
<tbody>
<tr>
<td>Renal Cell Carcinoma Cancer</td>
<td>Drug: pazopanib</td>
<td>Phase 3</td>
</tr>
<tr>
<td></td>
<td>Drug: placebo</td>
<td></td>
</tr>
</tbody>
</table>

Study Type: Interventional

Study Design:
- Allocation: Randomized
- Endpoint Classification: Safety/Efficacy Study
- Intervention Model: Parallel Assignment
- Masking: Double Blind (Subject, Caregiver, Investigator, Outcomes Assessor)
- Primary Purpose: Treatment

Official Title:
A Randomized, Double-blind, Placebo-controlled Phase III Study to Evaluate the Efficacy and Safety of Pazopanib as Adjuvant Therapy for Subjects With Localized or Locally Advanced RCC Following Nephrectomy

Resource links provided by NLM:
- [MedlinePlus](http://www.nlm.nih.gov/medlineplus) related topics: Cancer, Kidney Cancer
- [Drug Information](http://www.drugs.com) available for: Pazopanib
- [U.S. FDA Resources](http://www.fda.gov)

Further study details as provided by GlaxoSmithKline:

Primary Outcome Measures:
- Disease-free survival [Time Frame: approximately 4.5 years] 
  [Designated as safety issue: No]

Secondary Outcome Measures:
- Overall survival [Time Frame: approximately 9 years] 
  [Designated as safety issue: No]
- Disease-free survival rates at yearly time points (e.g., 1 year, 2 years, etc.). 
  [Time Frame: yearly for 4 or 5 years] [Designated as safety issue: No]
- Safety (frequency and severity of adverse events and laboratory abnormalities) [Time Frame: approximately 4.5 years] 
  [Designated as safety issue: Yes]
- Health Outcome (change from baseline in patients' self-reports on health outcome and quality of life as measured by two instruments: Cancer Therapy-Kidney Symptom Index-19 and EuroQOL-5D) 
  [Time Frame: approximately 4.5 years] [Designated as safety issue: No]

Estimated Enrollment: 1500
Study Start Date: November 2010
Estimated Study Completion Date: April 2017
Estimated Primary Completion Date: October 2015 (Final data collection date for primary outcome measure)

<table>
<thead>
<tr>
<th>Arms</th>
<th>Assigned Interventions</th>
</tr>
</thead>
<tbody>
<tr>
<td>Experimental: pazopanib</td>
<td>Drug: pazopanib</td>
</tr>
<tr>
<td>oral agent, administered at 800 mg daily (200 mg tablets x 4), continuously for 12 months. Dose can be reduced, interrupted or discontinued due to adverse events or intolerance</td>
<td></td>
</tr>
<tr>
<td>Placebo Comparator: placebo</td>
<td>Drug: placebo</td>
</tr>
<tr>
<td>placebo matching pazopanib 200 mg tablets, administered at 800 mg daily (200 mg tablets x 4), continuously for 12 months. Dose can be reduced, interrupted or discontinued due to adverse events or intolerance</td>
<td></td>
</tr>
</tbody>
</table>

Eligibility

Ages Eligible for Study: 18 Years and older
Genders Eligible for Study: Both
Accepts Healthy Volunteers: No

Criteria

Inclusion Criteria:
- Signed written informed consent
- Diagnosis of RCC with clear-cell or predominant clear-cell histology
- Subjects with non-metastatic disease (M0) fulfilling any of the following combinations of pathologic staging based on American Joint Committee on Cancer (AJCC) TNM staging version 2010 and Fuhrman nuclear grading:
  - pT2, G3 or G4, N0; or,
  - pT3, G any, N0; or,
  - pT4, G any, N0; or,
  - pT any, G any, N1
- Fulfill all of the following criteria of disease-free status at baseline:
  - Had complete gross surgical resection of all RCC via radical or partial
nephrectomy using either open or laparoscopic technique.
  o Baseline imaging of chest, abdomen and pelvis shows no metastasis or residual tumor lesions as confirmed centrally by an independent radiologist.

- Received no prior adjuvant or neo-adjuvant treatment for RCC
- Recovered from nephrectomy: any surgery related toxicities should be reduced to ≤ grade 1 per NCI Common Terminology Criteria for Adverse Events (CTCAE) (Version 4)
- Karnofsky performance scale (KPS) of ≥ 80
- Adequate organ system function

Exclusion Criteria:
- History of another malignancy. Exception: Subjects who have had another malignancy and have been disease-free for 5 years, or subjects with a history of completely resected non-melanomatus skin carcinoma or successfully treated in situ carcinoma are eligible
- Clinically significant gastrointestinal abnormalities that may increase the risk for gastrointestinal bleeding including, but not limited to:
  o Active peptic ulcer disease
  o Inflammatory bowel disease (e.g. ulcerative colitis, Crohn's disease), or other gastrointestinal conditions with increased risk of perforation
  o History of abdominal fistula, gastrointestinal perforation, or intra-abdominal abscess within 28 days prior to beginning study treatment
- Active diarrhea of any grade
- Clinically significant gastrointestinal abnormalities that may affect absorption of investigational product including, but not limited to:
  o Malabsorption syndrome
  o Major resection of the stomach or small bowel
- History of human immunodeficiency virus (HIV) infection
- History of active hepatitis
- Presence of uncontrolled infection.
- History of any one or more of the following cardiovascular conditions within the past 6 months:
  o Cardiac angioplasty or stenting
  o Myocardial infarction
  o Unstable angina
  o Coronary artery bypass graft surgery
  o Symptomatic peripheral vascular disease
- History of Class III or IV congestive heart failure, as defined by the New York Heart Association Classification of Congestive Heart Failure
- History of cerebrovascular accident including transient ischemic attack (TIA), pulmonary embolism or untreated deep venous thrombosis (DVT) within the past 6 months.
- Corrected QT interval (QTc) > 480 milliseconds (msec)
- Poorly controlled hypertension, defined as systolic blood pressure (SBP) of ≥140 mmHg or diastolic blood pressure (DBP) of ≥ 90mmHg.

Note: Initiation or adjustment of antihypertensive medication(s) is permitted prior to study entry. Blood pressure (BP) must be re-assessed on two occasions that are separated by a minimum of 1 hour; on each of these occasions, the mean (of 3 readings) SBP / DBP values from each BP assessment must be <140/90 mmHg in order for a subject to be eligible for the study (see Section 7.6.2 for instruction on blood pressure measurement and obtaining mean blood pressure values).

- Evidence of active bleeding or bleeding diathesis
- Any serious and/or unstable pre-existing medical, psychiatric, or other condition
A Study to Evaluate Pazopanib as an Adjuvant Treatment for Localized Renal Cell Ca...

that could interfere with subject's safety, provision of informed consent, or compliance to study procedures

- Unable or unwilling to discontinue use of prohibited medications for at least 14 days or five half-lives of a drug (whichever is longer) prior to the first dose of study treatment and for the duration of the study.
- Concurrent therapy given to treat cancer including treatment with an investigational agent or concurrent participation in another clinical trial involving anti-cancer investigational drug.
- Administration of an investigational drug within 30 days or 5 half-lives, whichever is longer, preceding the first dose of study treatment.
- Have a known immediate or delayed hypersensitivity reaction or idiosyncrasy to drugs chemically related to pazopanib or excipients that in the opinion of the investigator contraindicates their participation.
- Prior or current use of systemic anti-VEGF inhibitors, cytokines (e.g. interferon, interleukin 2).

**Contacts and Locations**

Please refer to this study by its ClinicalTrials.gov identifier: NCT01235962

**Contacts**

Contact: US GSK Clinical Call Center 877-379-3718  GSKClinicalSupportHD@gsk.com

**Show 295 Study Locations**

**Sponsors and Collaborators**

GlaxoSmithKline

**Investigators**

Study Director:  GSK Clinical Trials  GlaxoSmithKline

**More Information**

Publications:


**Responsible Party:** GlaxoSmithKline

**ClinicalTrials.gov Identifier:** NCT01235962  **History of Changes**

**Other Study ID Numbers:** 113387

**Study First Received:** October 14, 2010

**Last Updated:** July 12, 2012

**Health Authority:**

- Canada: Health Products and Foods Branch, Health Canada
- Spain: Agencia Espanola de Medicamentos y Productos Sanitarios
- Slovakia: State Institute for Drug Control
- Argentina: Ministry of Health - A.N.M.A.T
- Belgium: Agence Fédérale des Médicaments et des Produits de la Santé
- Brazil: National Health Surveillance Agency
- Italy: Agenzia Italiana del Farmaco
- Chile: Ministerio de Salud de Chile
- Poland: Office for Registration of Medicinal Products, Medical Devices and Biocidal Products
- Greece: National Organization of Medicines
- Hungary: National Institute of Pharmacy
- United Kingdom: Medicines and Healthcare Products Regulatory Authority

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