### Purpose

This pivotal phase III trial (CINC424B2301) is designed to compare the efficacy and safety of INC424 to Best Available Therapy (BAT) in subjects with polycythemia vera (PV) who are resistant to or intolerant of hydroxyurea (HU).

<table>
<thead>
<tr>
<th>Condition</th>
<th>Intervention</th>
<th>Phase</th>
</tr>
</thead>
<tbody>
<tr>
<td>Polycythemia Vera</td>
<td>Drug: INC424 tablets Other: Best Available Therapy (BAT)</td>
<td>Phase 3</td>
</tr>
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</table>

#### Study Details

- **Study Type:** Interventional
- **Study Design:** Allocation: Randomized Intervention Model: Parallel Assignment Masking: Open Label Primary Purpose: Treatment
- **Official Title:** Randomized, Open Label, Multicenter Phase III Study of Efficacy and Safety in Polycythemia Vera Subjects Who Are Resistant to or Intolerant of Hydroxyurea: JAK Inhibitor INC424 Tablets Versus Best Available Care

#### Resource links provided by NLM:

- Genetics Home Reference related topics: polycythemia vera
- Drug Information available for: Hydroxyurea
- U.S. FDA Resources

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**Further study details as provided by Incyte Corporation:**

**Primary Outcome Measures:**
- To compare the efficacy of INC424 to Best Available Therapy as assessed by both the absence of phlebotomy eligibility and reduction in spleen volume. [Time Frame: 32 Weeks] [Designated as safety issue: No]

**Secondary Outcome Measures:**
- To compare the proportion of subjects randomized to INC424 vs. Best Available Therapy achieving both durable absence of phlebotomy eligibility and durable spleen volume reduction. [Time Frame: 48 Weeks] [Designated as safety issue: No]
- To compare the proportion of subjects randomized to INC424 vs. Best Available Therapy achieving complete hematological remission. [Time Frame: 32 Weeks] [Designated as safety issue: No]
- To determine the proportion of subjects achieving a durable spleen volume reduction [Time Frame: 48 Weeks] [Designated as safety issue: No]
- To estimate the proportion of subjects achieving a durable complete hematological remission [Time Frame: 48 Weeks] [Designated as safety issue: No]
To estimate the proportion of subjects achieving a durable phlebotomy independence [Time Frame: 48 Weeks]
[Designated as safety issue: No]

To estimate the duration of both the absence of phlebotomy eligibility and reduction in spleen volume [Time Frame: Through study completion]
[Designated as safety issue: No]

To determine the overall clinicohematologic response rate [Time Frame: 32 Weeks] [Designated as safety issue: No]

To estimate the proportion of subjects achieving a durable complete or partial clinicohematologic response [Time Frame: 48 Weeks]
[Designated as safety issue: No]

To estimate the duration of the overall clinicohematologic response [Time Frame: Through study completion] [Designated as safety issue: No]

To estimate the proportion of subjects achieving both durable absence of phlebotomy eligibility and durable spleen volume reduction 48 weeks after the response was initially documented [Time Frame: Through study completion] [Designated as safety issue: No]

Safety and tolerability of INC424 as measured by adverse events, laboratory assessments, physical examination, vital signs, and ECG measurements [Time Frame: Through study completion] [Designated as safety issue: No]

Estimated Enrollment: 200
Study Start Date: October 2010
Estimated Study Completion Date: March 2014
Estimated Primary Completion Date: August 2013 (Final data collection date for primary outcome measure)

Arms

<table>
<thead>
<tr>
<th>Arms</th>
<th>Assigned Interventions</th>
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</thead>
<tbody>
<tr>
<td>Experimental: INC424 tablets</td>
<td>Drug: INC424 tablets</td>
</tr>
<tr>
<td>Starting dose of 10 mg BID with individualized dose titration ranging from 5 mg QD to 25 mg BID based on safety and efficacy</td>
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Best Available Therapy

Best Available Therapy (BAT) will be selected by the Investigator for each subject. BAT may not include experimental agents (i.e. those not approved for the treatment of any indication) as well as a limited number of other selected drugs.

Other: Best Available Therapy (BAT)

Best Available Therapy (BAT) will be selected by the investigator for each subject. BAT may not include experimental agents (i.e. those not approved for the treatment of any indication) as well as a limited number of other selected drugs.

Other Names:
- BAT:
  - Hydroxyurea
  - IFN/PEG-IFN
  - Pipobroman
  - Anagrelide
  - IMIDs
  - Observation only

Eligibility

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

Criteria

Inclusion Criteria:
- Subjects diagnosed with PV for at least 24 weeks prior to screening according to the 2008 World Health Organization criteria
- Subjects resistant to or intolerant of hydroxyurea
- Subjects with a phlebotomy requirement
- Subjects with a palpable splenomegaly and a spleen volume of greater than or equal to 450 cubic centimeters
- Subjects with an ECOG performance status of 0, 1 or 2

Exclusion Criteria:
- Women who are pregnant or nursing
- Subjects with inadequate liver or renal function
- Subjects with significant bacterial, fungal, parasitic, or viral infection requiring treatment
- Subjects with an active malignancy within the past 5 years, excluding specific skin cancers
- Subjects with known active hepatitis or HIV positivity
- Subjects who have previously received treatment with a JAK inhibitor
- Subjects being treated with any investigational agent

Contacts and Locations

http://clinicaltrials.gov/ct2/show/NCT01243944?term=CINC424B230...
Study of Efficacy and Safety in Polycythemia Vera Subjects Who Are Re... 

<table>
<thead>
<tr>
<th>Sponsors and Collaborators</th>
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</thead>
<tbody>
<tr>
<td>Incyte Corporation</td>
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<td>Novartis Pharmaceuticals</td>
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<tr>
<th>Investigators</th>
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<tbody>
<tr>
<td>Study Director: Srdan Verstovsek, MD, PhD, M.D. Anderson Cancer Center</td>
</tr>
<tr>
<td>Study Director: Bijoyesh Mookerjee, MD, Incyte Corporation</td>
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**More Information**

**Additional Information:**

- **Related Info:**
  - [Show 156 Study Locations](#)

**No publications provided**

- **Responsible Party:** Incyte Corporation
- **ClinicalTrials.gov Identifier:** NCT01243944
- **History of Changes**
- **Other Study ID Numbers:** CINC424B2301
- **Study First Received:** November 17, 2010
- **Last Updated:** January 29, 2013
- **Health Authority:** United States: Food and Drug Administration

**Keywords provided by Incyte Corporation:**
- INCB018424

**Additional relevant MeSH terms:**

- Polycythemia
- Polycythemia Vera
- Hematologic Diseases
- Myeloproliferative Disorders
- Bone Marrow Diseases
- Hydroxyurea
- Antineoplastic Agents
- Therapeutic Uses
- Pharmacologic Actions
- Antisickling Agents
- Hematologic Agents
- Enzyme Inhibitors
- Molecular Mechanisms of Pharmacological Action
- Nucleic Acid Synthesis Inhibitors

**ClinicalTrials.gov processed this record on April 01, 2013**