Jakafi® (ruxolitinib) Is the First and Only FDA-Approved Treatment for Steroid-Refractory Acute GVHD

**Indication and Usage**
Jakafi is indicated for treatment of steroid-refractory acute graft-versus-host disease (GVHD) in adult and pediatric patients 12 years and older.

**REACH1: An Open-Label, Single-Arm, Multicenter Study of Jakafi in Combination with Steroids**
Patients had Grade II-IV steroid-refractory acute GVHD (aGVHD)* occurring after allogeneic hematopoietic stem cell transplant; 71 patients were enrolled, of whom 49 were refractory to steroids alone and evaluable for efficacy.

- **73%** had Grade III or IV aGVHD†
- **84%** had visceral disease†
  — Includes patients with upper and lower GI and liver involvement‡

*Defined using Mount Sinai Acute GVHD International Consortium (MAGIC) criteria.

**Day 28 Responses Were Achieved in the Majority of Patients Treated With Jakafi**

**Primary Endpoint: ORR at Day 28**

<table>
<thead>
<tr>
<th>Patients, %</th>
<th>CR</th>
<th>VGPR</th>
<th>PR</th>
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</thead>
<tbody>
<tr>
<td>N = 49</td>
<td>57% (95% CI: 42, 71)</td>
<td>22%</td>
<td>4%</td>
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<tr>
<td>N = 49</td>
<td>31%</td>
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</table>

In the REACH1 trial, ORR was defined as complete response, very good partial response, or partial response by The Center for International Blood and Marrow Transplant Research (CIBMTR) criteria.

**Important Safety Considerations**
- Treatment with Jakafi can cause thrombocytopenia, anemia, and neutropenia. Perform a pre-treatment complete blood count (CBC) and monitor CBCs every 2 to 4 weeks until doses are stabilized, and then as clinically indicated. In patients with cytopenias, consider dose reductions, temporarily interrupting Jakafi, or transfusions, as clinically indicated.
- Serious bacterial, mycobacterial, fungal, and viral infections have occurred. Delay starting Jakafi until active serious infections have resolved. Evaluate patients prior to and while receiving Jakafi for risk factors and signs and symptoms of infection, and manage promptly. Use active surveillance and prophylactic antibiotics according to clinical guidelines.

Please see Important Safety Information for Jakafi on back for related and other risks.

**Indication and Usage**

Jakafi is indicated for treatment of steroid-refractory acute graft-versus-host disease (GVHD) in adult and pediatric patients 12 years and older.

**Important Safety Information**

- Treatment with Jakafi® (ruxolitinib) can cause thrombocytopenia, anemia and neutropenia, which are each dose-related effects. Perform a pre-treatment complete blood count (CBC) and monitor CBCs every 2 to 4 weeks until doses are stabilized, and then as clinically indicated.
- Manage thrombocytopenia by reducing the dose or temporarily interrupting Jakafi. Platelet transfusions may be necessary.
- Patients developing anemia may require blood transfusions and/or dose modifications of Jakafi.
- Severe neutropenia (ANC <0.5 × 10^9/L) was generally reversible by withholding Jakafi until recovery.
- Serious bacterial, mycobacterial, fungal and viral infections have occurred. Delay starting Jakafi until active serious infections have resolved. Observe patients receiving Jakafi for signs and symptoms of infection and manage promptly. Use active surveillance and prophylactic antibiotics according to clinical guidelines.
- Tuberculosis (TB) infection has been reported. Observe patients taking Jakafi for signs and symptoms of active TB and manage promptly. Prior to initiating Jakafi, evaluate patients for TB risk factors and test those at higher risk for latent infection. Consult a physician with expertise in the treatment of TB before starting Jakafi in patients with evidence of active or latent TB. Continuation of Jakafi during treatment of active TB should be based on the overall risk-benefit determination.
- Progressive multifocal leukoencephalopathy (PML) has occurred with Jakafi treatment. If PML is suspected, stop Jakafi and evaluate.
- Advise patients about early signs and symptoms of herpes zoster and to seek early treatment.
- Increases in hepatitis B viral load with or without associated elevations in alanine aminotransferase and aspartate aminotransferase have been reported in patients with chronic hepatitis B virus (HBV) infections. Monitor and treat patients with chronic HBV infection according to clinical guidelines.
- When discontinuing Jakafi, myeloproliferative neoplasm-related symptoms may return within one week. After discontinuation, some patients with myelofibrosis have experienced fever, respiratory distress, hypotension, DIC, or multi-organ failure. If any of these occur after discontinuation or while tapering Jakafi, evaluate and treat any intercurrent illness and consider restarting or increasing the dose of Jakafi. Instruct patients not to interrupt or discontinue Jakafi without consulting their physician. When discontinuing or interrupting Jakafi for reasons other than thrombocytopenia or neutropenia, consider gradual tapering rather than abrupt discontinuation.
- Non-melanoma skin cancers including basal cell, squamous cell, and Merkel cell carcinoma have occurred. Perform periodic skin examinations.
- Treatment with Jakafi has been associated with increases in total cholesterol, low-density lipoprotein cholesterol, and triglycerides. Assess lipid parameters 8-12 weeks after initiating Jakafi. Monitor and treat according to clinical guidelines for the management of hyperlipidemia.
- In myelofibrosis and polycythemia vera, the three most common nonhematologic adverse reactions (incidence >10%) were bruising, dizziness and headache. In acute graft-versus-host disease, the most common nonhematologic adverse reactions (incidence >50%) were infections and edema.
- Dose modifications may be required when administering Jakafi with strong CYP3A4 inhibitors or fluconazole or in patients with renal or hepatic impairment. Patients should be closely monitored and the dose titrated based on safety and efficacy.
- Use of Jakafi during pregnancy is not recommended and should only be used if the potential benefit justifies the potential risk to the fetus. Women taking Jakafi should not breastfeed during treatment and for two weeks after the final dose.

**Please see accompanying Full Prescribing Information for Jakafi.**

**References:**