Positive Quality Intervention: Ropeginterferon alfa-2b-njft (BESREMi) Use in Polycythemia Vera

**Description:** The purpose of this PQI is to discuss the use of ropeginterferon alfa-2b-njft (BESREMi) in the management of polycythemia vera (PV).

**Background:** Ropeginterferon alfa-2b-njft is indicated for the treatment of adults with polycythemia vera. It was approved by the FDA in 2021 for PV regardless of treatment history and the first interferon therapy specifically approved for PV. NCCN has designated ropeginterferon alfa-2b-njft as a preferred therapy option for high and symptomatic low-risk PV needing cytoreductive treatment. It is a subcutaneous injection given by a health care professional or by patient/caregiver, administered once every two weeks with the potential to change to every four weeks dosing after achievement of hematological stability for at least 1 year on a stable dose, offering patients a reduction in pill burden. The CONTINUATION-PV study found that 71% of patients achieved complete hematologic response (CHR) on ropeginterferon alfa-2b-njft at 36 months, compared to 51% of patients achieving CHR on hydroxyurea. Ropeginterferon alfa-2b-njft also achieved CHR in 80% (based only on lab parameters) with all three counts (hematocrit, platelets, and WBCs) over the 7 year observation period in the majority of patients, according to the PEGINVERA study. Ropeginterferon alfa-2b-njft also showed similar rates of adverse reactions compared to hydroxyurea, with both arms sharing the incidence rate of thromboembolic adverse events of 1.2% per patient year.

**PQI Process:**

- Confirm diagnosis for polycythemia vera
- Assess the patient’s need for ropeginterferon alfa-2b-njft versus other cytoreductive options (i.e., hydroxyurea, peginterferon)
- Assess the patient’s medical history for contraindications to ropeginterferon alfa-2b-njft
  - Hypersensitivity to interferons, including interferon alfa-2b, or any component of the formulation; existence of, or history of severe psychiatric disorders, particularly severe depression, suicidal ideation, or suicide attempt; moderate (Child-Pugh class B) or severe (Child-Pugh class C) hepatic impairment; history or presence of active serious or untreated autoimmune disease; immunosuppressed transplant recipients
  - Avoid in patients with uncontrolled hypertension, congestive heart failure, arrhythmias, significant arterial stenosis, unstable angina, recent stroke or myocardial infarction
- Assess pregnancy status in females of reproductive age
- Review the patient’s medication history for use of hydroxyurea in anticipation of possible need for ropeginterferon alfa-2b-njft dosage adjustment as detailed below
- Monitoring
  - Perform blood counts at baseline, every 2 weeks during titration and at least every 3-6 months during maintenance treatment
  - Eye exams at baseline and during treatment
  - Monitor for signs of suicidal ideation
  - Monitor cardiovascular toxicity
  - Discontinue if any of the following occur: medically unmanageable endocrine toxicities, severe cardiac disease, hypersensitivity reactions, pancreatitis, colitis, pulmonary function impairment
- Adverse Events
  - Boxed warning: Risk of Serious Disorders: Interferon alfa products may cause or aggravate fatal or life-threatening neuropsychiatric, autoimmune, ischemic, and infectious disorders
  - Most common Adverse Events: Flu like symptoms, itching, sore throat, hypersensitivity

**IMPORTANT NOTICE:** NCODA has developed this Positive Quality Intervention platform. This platform is intended as an educational aid, does not provide individual medical advice, and does not substitute for the advice of a qualified healthcare professional. This platform does not cover all existing information related to the possible uses, directions, doses, precautions, warning, interactions, adverse effects, or risks associated with the medication. The materials contained in this platform do not constitute or imply endorsement, recommendation, or favoring of this medication by NCODA. NCODA does not ensure the accuracy of the information presented and assumes no liability relating to its accuracy. All decisions related to taking this medication should be made with the guidance and under the direction of a qualified healthcare professional. It is the individual’s sole responsibility to seek guidance from a qualified healthcare professional. Updated 5.10.24
reactions, ophthalmologic toxicity, hepatotoxicity, endocrine toxicity, depression and suicide

- **Dose Modification**

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<thead>
<tr>
<th></th>
<th>Starting Dose</th>
<th>Increasing Dose</th>
<th>Maximum Dose</th>
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<tbody>
<tr>
<td>Transitioning from</td>
<td>50 mcg SQ</td>
<td>50 mcg SQ every 2 weeks until hematological parameters stabilize (hematocrit &lt;45%, platelets &lt;400,000/mm³, and leukocytes &lt;10,000/mm³)</td>
<td>500 mcg</td>
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<tr>
<td>Hydroxyurea (HU)</td>
<td>every 2</td>
<td>Gradually taper the HU by reducing the total biweekly hydroxyurea dose by 20% to 40%</td>
<td>every 2</td>
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<tr>
<td></td>
<td>weeks in combo</td>
<td>every 2 weeks during weeks 3-12; discontinue hydroxyurea by week 13</td>
<td>weeks</td>
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<tr>
<td>Not on Hydroxyurea</td>
<td>100 mcg</td>
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<td>every 2</td>
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- Patients with impaired hepatic function: no adjustment; use is contraindicated in Child-Pugh class B/C
- Patients with impaired renal function: avoid use in patients with eGFR <30 mL/minute or if renal impairment occurs during treatment
- Patients who experience adverse reactions: dosage adjustments and discontinuation recommendations for LFT elevations, cytopenia, and depression are listed in the **package insert**

**Patient-Centered Activities:**¹

- **Patient Education**
  - Advise patients to read the FDA-approved medication guide and instructions for use
  - Ensure eye exams and labs are scheduled prior to initiating treatment and as clinically necessary
  - Educate on good oral hygiene and regular dental examination
  - Counsel on proper storage, preparation, and administration techniques if administering at home
  - Instruct patients to report any signs of an allergic reaction, like rash, hives, itching, red, swollen, blistered or peeling skin with or without fever, wheezing, tightness in the chest or throat, trouble breathing, or swelling of the mouth, face, lips, tongue, or throat
  - Direct patients to report any signs of depression, suicidal thoughts, or abnormal thinking
  - Inform patients to report symptoms of tiredness, frequent urination, and increase in thirst
  - Advise females of reproductive age to use an effective method of contraception during treatment and for at least 8 weeks after the final dose
  - Advise women not to breastfeed during treatment and for 8 weeks after the final dose
  - Ensure the patient is on aspirin 81-100 mg daily

**References:**

1. BESREMI (ropeginterferon alfa-2b-njft) package insert.