Daurismo

**Description**

Daurismo (glasdegib)

**Background**

Daurismo (glasdegib) is an inhibitor of the Hedgehog pathway. Daurismo binds to and inhibits Smoothened, a transmembrane protein involved in hedgehog signal transduction. Daurismo in combination with low-dose cytarabine inhibits increases in tumor size and reduces the percentage of CD45+/CD33+ blasts in the marrow (1).

**Regulatory Status**

FDA-approved indication: Daurismo is a Hedgehog pathway inhibitor indicated, in combination with low-dose cytarabine, for the treatment of newly-diagnosed acute myeloid leukemia (AML) in adult patients who are ≥75 years old or who have comorbidities that preclude use of intensive induction chemotherapy (1).

Limitations of use: Daurismo has not been studied in patients with the comorbidities of severe renal impairment or moderate-to-severe hepatic impairment (1).

Daurismo has a boxed warning regarding embryo-fetal death or severe birth defects. Pregnancy testing should be done in females of reproductive potential prior to initiation of Daurismo treatment. Females of reproductive potential should be advised to use effective contraception during treatment with Daurismo and for at least 30 days after the last dose. Males with a pregnant partner or a female partner of reproductive potential should be advised to use condoms during treatment with Daurismo and for at least 30 days after the last dose (1).
Complete blood counts, electrolytes, renal, and hepatic function should be assessed prior to initiation of Daurismo and at least once weekly for the first month. Electrolytes and renal function should be monitored once monthly for the duration of therapy. Serum creatinine kinase levels should be obtained prior to initiating Daurismo and as indicated clinically thereafter. Electrocardiograms (ECGs) should be monitored prior to initiation of Daurismo, approximately one week after initiation, and then once monthly for the next two months to assess for QTc prolongation (1).

The safety and effectiveness of Daurismo in pediatric patients have not been established (1).

Related policies

Policy

This policy statement applies to clinical review performed for pre-service (Prior Approval, Precertification, Advanced Benefit Determination, etc.) and/or post-service claims.

Daurismo may be considered medically necessary in patients 18 years of age or older with acute myeloid leukemia (AML) and if the conditions indicated below are met.

Daurismo is considered investigational in patients less than 18 years of age and for all other indications.

Prior-Approval Requirements

Age

18 years of age or older

Diagnosis

Patient must have the following:

Newly-diagnosed acute myeloid leukemia (AML)

AND ALL of the following:

1. Used in combination with low-dose cytarabine
2. Patient is 75 years of age or older OR patient has comorbidities that preclude the use of intensive induction chemotherapy
3. Prescriber agrees to monitor electrocardiograms (ECGs) for QTc prolongation
4. Prescriber agrees to advise females of reproductive potential to use effective
contraception during treatment and for at least 30 days after the last dose
5. Prescriber agrees to advise males with female partners of reproductive potential to use condoms during treatment and for at least 30 days after the last dose

Prior – Approval Renewal Requirements

Age
18 years of age or older

Diagnosis

Patient must have the following:

Acute myeloid leukemia (AML)

AND ALL of the following:
1. Used in combination with low-dose cytarabine
2. NO disease progression or unacceptable toxicity
3. Prescriber agrees to monitor electrocardiograms (ECGs) for QTc prolongation
4. Prescriber agrees to advise females of reproductive potential to use effective contraception during treatment and for at least 30 days after the last dose
5. Prescriber agrees to advise males with female partners of reproductive potential to use condoms during treatment and for at least 30 days after the last dose

Policy Guidelines

Pre - PA Allowance
None

Prior - Approval Limits

Quantity

<table>
<thead>
<tr>
<th>Strength</th>
<th>Quantity Limit per 90 days</th>
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<tbody>
<tr>
<td>25 mg tablets</td>
<td>180 tablets per 90 days <strong>OR</strong></td>
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<td>100 mg tablets</td>
<td>90 tablets per 90 days</td>
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**Duration**

12 months

**Prior – Approval Renewal Limits**

Same as above

### Rationale

**Summary**

Daurismo (glasdegib) is an inhibitor of the Hedgehog pathway. Daurismo binds to and inhibits Smoothened, a transmembrane protein involved in hedgehog signal transduction. Daurismo in combination with low-dose cytarabine inhibits increases in tumor size and reduces the percentage of CD45+/CD33+ blasts in the marrow. The safety and effectiveness of Daurismo in pediatric patients have not been established (1).

Prior authorization is required to ensure the safe, clinically appropriate and cost-effective use of Daurismo while maintaining optimal therapeutic outcomes.

**References**


### Policy History

<table>
<thead>
<tr>
<th>Date</th>
<th>Action</th>
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<tbody>
<tr>
<td>December 2018</td>
<td>Addition to PA</td>
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<tr>
<td>March 2019</td>
<td>Annual review</td>
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### Keywords

This policy was approved by the FEP® Pharmacy and Medical Policy Committee on March 15, 2019 and is effective on April 1, 2019.