



Gene Therapies for Sickle Cell Disease

Draft Questions for Deliberation and Voting: July 27th, 2023 Public Meeting

These questions are intended for the deliberation of the CTAF voting body at the public meeting.

Patient population for all questions: Adolescents and adults with severe sickle cell disease (SCD) who do not have access to, or cannot receive, hematopoietic stem cell transplantation (HSCT) from a matched sibling or haploidentical donor.

Clinical Evidence

1. Given the currently available evidence, is the evidence adequate to demonstrate that the net health benefit of exagamglogene autotemcel (exa-cel) is superior to that provided by standard of care (i.e., hydroxyurea, chronic blood transfusions, pain medication, iron chelation)?

Yes No

2. Given the currently available evidence, is the evidence adequate to demonstrate that the net health benefit of lovo-cel is superior to that provided by standard of care (i.e., hydroxyurea, chronic blood transfusions, pain medication, iron chelation)?

Yes No

3. Given the currently available evidence, is the evidence adequate to distinguish the net health benefit between exa-cel and lovo-cel?

Yes No

If yes, answer question 3a:

- 3a. Given the currently available evidence, which product is superior based on its net health benefit?

Exa-cel Lovo-cel

Contextual Considerations and Potential Other Benefits or Disadvantages

Please vote on the following contextual considerations:

When making judgments of overall long-term value for money, what is the relative priority that should be given to any effective treatment for SCD, on the basis of the following contextual considerations:

1=Very low priority; 2=Low priority; 3=Average priority; 4=High priority; 5=Very high priority

4. Acuity of need for treatment of individual patients based on short-term risk of death or progression to permanent disability
5. Magnitude of the lifetime impact on individual patients of the condition being treated

Please vote on the following potential other benefits or disadvantages:

What are the relative effects of exa-cel* versus standard of care on the following outcomes that inform judgment of the overall long-term value for money of exa-cel*?

1=Major negative effect; 2=Minor negative effect; 3=No difference; 4=Minor positive effect; 5=Major positive effect

6. Patients' ability to achieve major life goals related to education, work, or family life
7. Caregivers' quality of life and/or ability to achieve major life goals related to education, work, or family life
8. Patients' ability to manage and sustain treatment given the complexity of regimen
9. Society's goal of reducing health inequities

**Note: if the answer to question 3 is No, then only one vote will be taken, with the question combining the relative effects of exa-cel and lovo-cel on these outcomes.*

What are the relative effects of lovo-cel versus standard of care on the following outcomes that inform judgment of the overall long-term value for money of lovo-cel?

1=Major negative effect; 2=Minor negative effect; 3=No difference; 4=Minor positive effect; 5=Major positive effect

10. Patients' ability to achieve major life goals related to education, work, or family life
11. Caregivers' quality of life and/or ability to achieve major life goals related to education, work, or family life
12. Patients' ability to manage and sustain treatment given the complexity of regimen
13. Society's goal of reducing health inequities

Long-Term Value for Money

14. Given the available evidence on comparative effectiveness and incremental cost-effectiveness, and considering other benefits, disadvantages, and contextual considerations, what is the long-term value for money of treatment at assumed pricing with exa-cel versus standard of care?*

 - a. Low long-term value for money at assumed pricing
 - b. Intermediate long-term value for money at assumed pricing
 - c. High long-term value for money at assumed pricing

15. Given the available evidence on comparative effectiveness and incremental cost-effectiveness, and considering other benefits, disadvantages, and contextual considerations, what is the long-term value for money of treatment at current pricing with lovo-cel versus standard of care?†

 - a. Low long-term value for money at assumed pricing
 - b. Intermediate long-term value for money at assumed pricing
 - c. High long-term value for money at assumed pricing

**This vote will only be taken if the price becomes available for exa-cel.*

†This vote will only be taken if the price becomes available for lovo-cel.