

Medical Policy Manual **Approved New: Do Not Implement until 12/31/24**

Donanemab-azbt (KISUNLA™)

IMPORTANT REMINDER

We develop Medical Policies to provide guidance to Members and Providers. This Medical Policy relates only to the services or supplies described in it. The existence of a Medical Policy is not an authorization, certification, explanation of benefits or a contract for the service (or supply) that is referenced in the Medical Policy. For a determination of the benefits that a Member is entitled to receive under his or her health plan, the Member's health plan must be reviewed. If there is a conflict between the medical policy and a health plan or government program (e.g., TennCare), the express terms of the health plan or government program will govern.

POLICY

I. INDICATIONS

The indications below including FDA-approved indications and compendial uses are considered a covered benefit provided that all the approval criteria are met and the member has no exclusions to the prescribed therapy.

FDA-Approved Indication

Kisunla is indicated for the treatment of Alzheimer's disease. Treatment with Kisunla should be initiated in patients with mild cognitive impairment or mild dementia stage of disease, the population in which treatment was initiated in the clinical trials.

All other indications are considered experimental/investigational and not medically necessary.

II. DOCUMENTATION

Submission of the following information is necessary to initiate the prior authorization review:

A. Initial Requests:

1. Genetic testing results documenting a mutation in amyloid precursor protein (*APP*), presenilin-1 (*PSEN1*), or presenilin-2 (*PSEN2*), if applicable.
2. Clinical documentation to support early onset Alzheimer's Disease, if applicable.
3. Medical records (e.g., chart notes) documenting the following:
 - i. Diagnosis of mild cognitive impairment due to Alzheimer's Disease or mild Alzheimer's Disease.
 - ii. Baseline assessments for any of the following assessment tools:
 - a. Clinical Dementia Rating-Global Score (CDR-GS)
 - b. Mini-Mental Status Examination (MMSE)
 - c. Montreal Cognitive Assessment (MoCA)
4. Presence of amyloid pathology documented by either of the following:
 - i. Baseline positron emission tomography (PET) scan
 - ii. Lumbar puncture results
5. Recent (within one year) brain magnetic resonance imaging (MRI) prior to initiating treatment.

B. Continuation requests (where applicable):

1. Medical records (e.g., chart notes) documenting the most recent (less than 1 month prior to continuation request) assessment tool for any of the following:
 - i. Clinical Dementia Rating-Global Score (CDR-GS)
 - ii. Mini-Mental Status Exam (MMSE)
 - iii. Montreal Cognitive Assessment (MoCA)



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2. Brain magnetic resonance imaging (MRI) results prior to the 2nd dose, 3rd dose, 4th dose, and 7th dose.

III. EXCLUSIONS

Coverage will not be provided for members with any of the following conditions:

- A. Suspected neurodegenerative etiology of cognitive impairment other than Alzheimer's disease (AD), including but not limited to frontotemporal lobar degeneration (FTLD) or Lewy body disease (i.e., meeting consensus criteria for possible or probable dementia with Lewy bodies).
- B. History of transient ischemic attacks (TIA), stroke, or seizures within the past 12 months.
- C. Bleeding disorder that is not under adequate control (including a platelet count <50,000 or international normalized ratio [INR] >1.5).
- D. Kisunla will not be used in combination with any other amyloid beta-directed antibodies (e.g., aducanumab, lecanemab).

IV. PRESCRIBER SPECIALTIES

This medication must be prescribed by or in consultation with a geriatrician, neurologist, psychiatrist, or neuropsychiatrist.

V. CRITERIA FOR INITIAL APPROVAL

Alzheimer's Disease

Authorization of 7 months may be granted for treatment of Alzheimer's Disease (AD) when all of the following criteria are met:

- A. Member must meet one of the following criteria:
 1. Member is 50 years of age or older
 2. If less than 50 years of age, member has a genetic mutation in amyloid precursor protein (*APP*), presenilin-1 (*PSEN1*), or presenilin-2 (*PSEN2*), or other clinical documentation to support early onset AD.
- B. Member must have mild cognitive impairment due to AD or mild AD dementia.
- C. Member must have objective evidence of cognitive impairment at baseline (Appendix A).
- D. Member must have one of the following scores at baseline on any of the following assessment tools:
 1. Clinical Dementia Rating-Global Score (CDR-GS) of 0.5 or 1 (Appendix B).
 2. Mini-Mental Status Examination (MMSE) score of 21 – 30 (Appendix C).
 3. Montreal Cognitive Assessment (MoCA) score of greater than or equal to 16 (Appendix D).
- E. Member must meet one of the following criteria:
 1. Have a positron emission tomography (PET) scan confirming the presence of amyloid pathology.
 2. Have results from a lumbar puncture confirming at least one of the following:
 - i. Presence of elevated phosphorylated tau (P-tau) protein and/or elevated total tau (T-tau) protein, and reduced beta amyloid-42 (AB42).
 - ii. Low AB42/AB40 ratio
 - iii. Elevated P-Tau/AB42 ratio
 - iv. Elevated T-tau/AB42 ratio
- F. Member must have a recent brain magnetic resonance imaging (MRI) within one year prior to initiating treatment to evaluate for pre-existing Amyloid Related Imaging Abnormalities (ARIA).
- G. Member meets one of the following regarding apolipoprotein E ϵ 4 (ApoE ϵ 4) status:



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1. Genotype testing for ApoE ϵ 4 status has been performed prior to initiation of treatment to inform member of the risk of developing ARIA.
2. Genotype testing has not been performed and the prescriber has informed the member that it cannot be determined if they are ApoE ϵ 4 homozygous and may be at higher risk for ARIA.
- H. If there is concurrent use of antithrombotic medications (aspirin, other antiplatelets, or anticoagulants), the member has been on a stable dose for at least 4 weeks prior to initiation of the requested medication.
- I. Member and/or provider must currently be participating in a provider-enrolled patient registry that collects information on treatments for Alzheimer's disease (e.g., Alzheimer's Network for Treatment and Diagnostics (ALZ-NET)).

VI. CONTINUATION OF THERAPY

Authorization of 12 months (first reauthorization after the initial 7-month approval period) may be granted for members requesting continuation of therapy when all of the following criteria are met:

- A. Member has met all initial authorization criteria at the time of initial approval.
- B. Member has been evaluated for evidence of amyloid-related imaging abnormalities (ARIA) on MRI prior to the 2nd dose, 3rd dose, 4th dose, and 7th dose (Appendix E).
 1. For members with radiographic evidence of ARIA-E:
 - i. Dosing may continue based on clinical judgement, if applicable, for members that meet the following criteria:
 - a. Member has mild ARIA-E on MRI and is asymptomatic or has mild clinical symptoms
 - ii. Dosing should be suspended until MRI demonstrates radiographic resolution and symptoms resolve for members that meet any of the following criteria:
 - a. Member has mild ARIA-E on MRI and has moderate or severe clinical symptoms
 - b. Member has moderate ARIA-E on MRI and is asymptomatic or has mild, moderate, or severe clinical symptoms
 - c. Member has severe ARIA-E on MRI and is asymptomatic or has mild, moderate, or severe clinical symptoms
 2. For members with radiographic evidence of ARIA-H:
 - i. Dosing may continue for members that meet the following criteria:
 - a. Member has mild ARIA-H on MRI and is asymptomatic
 - ii. Dosing should be suspended until MRI demonstrates radiographic stabilization and symptoms resolve for members that meet any of the following criteria:
 - a. Member has mild ARIA-H on MRI and is symptomatic
 - b. Member has moderate ARIA-H on MRI and is asymptomatic or symptomatic
 - c. Member has severe ARIA-H on MRI and is asymptomatic or symptomatic
- C. Member and/or provider continues to participate in a provider-enrolled patient registry that collects information on treatments for Alzheimer's disease (e.g., Alzheimer's Network for Treatment and Diagnostics (ALZ-NET)).

Authorization of 12 months (reauthorizations beyond initial 19 months of therapy) may be granted for members requesting continuation of therapy when all of the following criteria are met:

- A. Member has met all initial authorization criteria at the time of initial approval.
- B. Member has a positive clinical response as evidenced by stabilization or slowing of disease progression as documented by any of the following (Note: repeat assessment tool(s) must be the same tool that was submitted upon initial request):
 1. CDR-Global Score (i.e., score of 0.5 or 1)

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2. MMSE (i.e., decline of 3 points or less per year)
3. MoCA (i.e., score of greater than or equal to 16)

Note: Continuation requests for members with assessment scores outside of the provided ranges (i.e., mild dementia) or who have progressed greater than the provided rate of decline may be reviewed on a case-by-case basis.

- C. Member and/or provider continues to participate in a provider-enrolled patient registry that collects information on treatments for Alzheimer's disease (e.g., Alzheimer's Network for Treatment and Diagnostics (ALZ-NET)).

VII. APPENDICES

Appendix A: Summary of clinical and cognitive evaluation for MCI due to AD

- Cognitive concern reflecting a change in cognition reported by patient or informant or clinician (i.e., historical or observed evidence of decline over time)
- Objective evidence of impairment in one or more cognitive domains, typically including memory (i.e., formal or bedside testing to establish level of cognitive function in multiple domains)
- Preservation of independence in functional abilities
- Not demented

Appendix B: Clinical Dementia Rating (CDR) Scale

The CDR is obtained through semi-structured interviews of patients and informants with cognitive functioning rated on a 5-point scale in the following domains: memory, orientation, judgment and problem solving, community affairs, home and hobbies, and personal care. The score relates to the member's level of dementia:

- 0 = Normal
- 0.5 = Very Mild Dementia
- 1 = Mild Dementia
- 2 = Moderate Dementia
- 3 = Severe Dementia

Appendix C: Mini-Mental Status Exam (MMSE)

The MMSE is scored on a 30-point scale, with items that assess orientation (temporal and spatial; 10 points), memory (registration and recall; 6 points), attention/concentration (5 points), language (verbal and written, 8 points), and visuospatial function (1 point). The score relates to the member's level of dementia:

- 25 – 30 suggests normal cognition
- 20 – 24 suggests mild dementia
- 13 – 20 suggests moderate dementia
- Less than 12 suggests severe dementia

Appendix D: Montreal Cognitive Assessment (MoCA)

Per MoCA assessment, average scores for the following ranges are:

- Mild Cognitive Impairment: 19 – 25
- Mild Dementia: 11 – 21
- Normal: 26 and above

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Appendix E: ARIA MRI Classification Criteria

ARIA Type	Radiographic Severity		
	Mild	Moderate	Severe
ARIA-E	FLAIR hyperintensity confined to sulcus and or cortex/subcortical white matter in one location < 5 cm	FLAIR hyperintensity 5 to 10 cm in single greatest dimension, or more than 1 site of involvement, each measuring < 10 cm	FLAIR hyperintensity > 10 cm with associated gyral swelling and sulcal effacement. One or more separate/independent sites of involvement may be noted.
ARIA-H microhemorrhage	≤ 4 new incident microhemorrhages	5 to 9 new incident microhemorrhages	10 or more new incident microhemorrhages
ARIA-H superficial siderosis	1 new* focal area of superficial siderosis	2 new focal areas of superficial siderosis	> 2 new areas of superficial siderosis

* includes new or worsening superficial siderosis

APPLICABLE TENNESSEE STATE MANDATE REQUIREMENTS

BlueCross BlueShield of Tennessee's Medical Policy complies with Tennessee Code Annotated Section 56-7-2352 regarding coverage of off-label indications of Food and Drug Administration (FDA) approved drugs when the off-label use is recognized in one of the statutorily recognized standard reference compendia or in the published peer-reviewed medical literature.

ADDITIONAL INFORMATION

For appropriate chemotherapy regimens, dosage information, contraindications, precautions, warnings, and monitoring information, please refer to one of the standard reference compendia (e.g., the NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines®) published by the National Comprehensive Cancer Network®, Drugdex Evaluations of Micromedex Solutions at Truven Health, or The American Hospital Formulary Service Drug Information).

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EFFECTIVE DATE

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