

Clinical Policy: Cariprazine (Vraylar)

Reference Number: CP.PMN.91

Effective Date: 11.16.16

Last Review Date: 02.20

Line of Business: Commercial, Medicaid

[Revision Log](#)

See [Important Reminder](#) at the end of this policy for important regulatory and legal information.

Description

Cariprazine (Vraylar[®]) is an atypical antipsychotic.

FDA Approved Indication(s)

Vraylar is indicated for:

- Treatment of schizophrenia in adults
- Acute treatment of manic or mixed episodes associated with bipolar I disorder in adults
- Treatment of depressive episodes associated with bipolar I disorder (bipolar depression) in adults

Policy/Criteria

Provider must submit documentation (such as office chart notes, lab results or other clinical information) supporting that member has met all approval criteria.

It is the policy of health plans affiliated with Centene Corporation[®] that Vraylar is **medically necessary** when the following criteria are met:

I. Initial Approval Criteria

A. Bipolar Disorder and Schizophrenia (must meet all):

1. Diagnosis of bipolar disorder or schizophrenia;
2. Age \geq 18 years;
3. Failure of two preferred atypical antipsychotics (e.g., aripiprazole, ziprasidone, quetiapine, risperidone, or olanzapine) at up to maximally indicated doses, each used for \geq 4 weeks, unless contraindicated or clinically significant adverse effects are experienced;
4. Dose does not exceed any of the following:
 - a. Schizophrenia or manic or mixed episodes of bipolar I disorder: 6 mg (1 capsule) per day;
 - b. Depressive episodes of bipolar I disorder: 3 mg (1 capsule) per day.

Approval duration:

Medicaid – 12 months

Commercial – Length of Benefit

B. Other diagnoses/indications

1. Refer to the off-label use policy for the relevant line of business if diagnosis is NOT specifically listed under section III (Diagnoses/Indications for which coverage is NOT authorized): CP.CPA.09 for commercial and CP.PMN.53 for Medicaid.

II. Continued Therapy

A. All Indications in Section I (must meet all):

1. Currently receiving medication via Centene benefit, or documentation supports that member is currently receiving Vraylar for bipolar disorder or schizophrenia and has received this medication for at least 30 days;
2. Member is responding positively to therapy;
3. If request is for a dose increase, new dose does not exceed any of the following:
 - a. Schizophrenia or manic or mixed episodes of bipolar I disorder: 6 mg (1 capsule) per day;
 - b. Depressive episodes of bipolar I disorder: 3 mg (1 capsule) per day.

Approval duration:

Medicaid – 12 months

Commercial – Length of Benefit

B. Other diagnoses/indications (must meet 1 or 2):

1. Currently receiving medication via Centene benefit and documentation supports positive response to therapy.

Approval duration: Duration of request or 12 months (whichever is less); or

2. Refer to the off-label use policy for the relevant line of business if diagnosis is NOT specifically listed under section III (Diagnoses/Indications for which coverage is NOT authorized): CP.CPA.09 for commercial and CP.PMN.53 for Medicaid.

III. Diagnoses/Indications for which coverage is NOT authorized:

- ### A. Non-FDA approved indications, which are not addressed in this policy, unless there is sufficient documentation of efficacy and safety according to the off label use policies – CP.CPA.09 for commercial and CP.PMN.53 for Medicaid or evidence of coverage documents.

IV. Appendices/General Information

Appendix A: Abbreviation/Acronym Key

FDA: Food and Drug Administration

Appendix B: Therapeutic Alternatives

This table provides a listing of preferred alternative therapy recommended in the approval criteria. The drugs listed here may not be a formulary agent for all relevant lines of business and may require prior authorization.

Drug Name	Dosing Regimen	Dose Limit/ Maximum Dose
aripiprazole (Abilify®)	Bipolar Disorder and Schizophrenia Adults: 10 to 15 mg PO QD	30 mg/day
olanzapine (Zyprexa®)	Schizophrenia Initial: 5 to 10 mg PO QD; target: 10 mg PO QD Bipolar Disorder	20 mg/day

Drug Name	Dosing Regimen	Dose Limit/ Maximum Dose
	Monotherapy: 10 to 15 mg PO QD; adjunct to lithium or valproate: 10 mg PO QD	
quetiapine (Seroquel [®])	Schizophrenia Initial: 25 mg PO BID; target: 400 to 800 mg/day Bipolar Disorder Initial: 50 mg PO BID; target: 400 to 800 mg/day	800 mg/day
risperidone (Risperdal [®])	Schizophrenia Initial: 1 mg PO BID or 2 mg PO QD; target: 4 to 8 mg PO QD Bipolar Disorder 2 to 3 mg PO QD	Schizophrenia: 16 mg/day Bipolar Disorder: 6 mg/day
ziprasidone (Geodon [®])	Schizophrenia 20 mg PO BID Bipolar Disorder Initial: 40 mg PO BID; target: 40 to 80 mg PO BID	160 mg/day

Therapeutic alternatives are listed as Brand name[®] (generic) when the drug is available by brand name only and generic (Brand name[®]) when the drug is available by both brand and generic.

Appendix C: Contraindications/Boxed Warnings

- Contraindication(s): known hypersensitivity to Vraylar
- Boxed warning(s): Elderly patients with dementia-related psychosis treated with antipsychotic drugs are at an increased risk of death. Vraylar is not approved for the treatment of patients with dementia-related psychosis.

V. Dosage and Administration

Indication	Dosing Regimen	Maximum Dose
Schizophrenia	1.5 mg to 6 mg PO QD	6 mg/day
Bipolar I disorder	Manic or mixed episodes: 3 mg to 6 mg PO QD Depressive episodes: 1.5 mg or 3 mg PO QD	Manic or mixed episodes: 6 mg/day Depressive episodes: 3 mg/day

VI. Product Availability

Capsules: 1.5 mg, 3 mg, 4.5 mg, 6 mg

VII. References

1. Vraylar Prescribing Information. Irvine, CA: Allergan USA, Inc.; May 2019. Available at: <http://www.vraylar.com/>. November 30, 2019.
2. Lehman AF, Lieberman JA, Dixon LB et al. Practice guideline for the treatment of patients with schizophrenia, second edition. Am J Psychiatry. 2004 Feb;161(2 Suppl):1-56.

3. American Psychiatric Association. Practice guideline for the treatment of patients with bipolar disorder. Am J Psychiatry. 2002 Apr;159(4 Suppl):1-50.
4. American Psychiatric Association Practice Guideline for the Treatment of Patients with Bipolar Disorder: Second Edition (2010). Available at: http://psychiatryonline.org/pb/assets/raw/sitewide/practice_guidelines/guidelines/bipolar.pdf. Accessed November 30, 2019.
5. Crismon ML, Argo TR, Bendele SD, et al. Texas Medication Algorithm Project Procedural Manual: Bipolar Disorder Algorithms. July 2007. Available at: https://www.jpshhealthnet.org/sites/default/files/tmap_bipolar_2007.pdf. Accessed November 30 2019.
6. Micromedex® Healthcare Series [Internet database]. Greenwood Village, Colo: Thomson Healthcare. Updated periodically. Accessed November 30, 2019.

Reviews, Revisions, and Approvals	Date	P&T Approval Date
1Q18 annual review: Policy generated from existing commercial policy – CP.CPA.221; No significant change from previously approved corporate policy; New for Medicaid; Age added for schizophrenia per safety guidance endorsed by Centene Medical Affairs.	11.13.17	02.18
1Q 2019 annual review: no significant changes; references reviewed and updated.	10.30.18	02.19
RT4: updated policy to add newly FDA-approved indication for depressive episodes associated with bipolar I disorder; references updated.	06.14.19	
1Q 2020 annual review: no significant changes; references reviewed and updated.	11.30.19	02.20

Important Reminder

This clinical policy has been developed by appropriately experienced and licensed health care professionals based on a review and consideration of currently available generally accepted standards of medical practice; peer-reviewed medical literature; government agency/program approval status; evidence-based guidelines and positions of leading national health professional organizations; views of physicians practicing in relevant clinical areas affected by this clinical policy; and other available clinical information. The Health Plan makes no representations and accepts no liability with respect to the content of any external information used or relied upon in developing this clinical policy. This clinical policy is consistent with standards of medical practice current at the time that this clinical policy was approved. “Health Plan” means a health plan that has adopted this clinical policy and that is operated or administered, in whole or in part, by Centene Management Company, LLC, or any of such health plan’s affiliates, as applicable.

The purpose of this clinical policy is to provide a guide to medical necessity, which is a component of the guidelines used to assist in making coverage decisions and administering benefits. It does not constitute a contract or guarantee regarding payment or results. Coverage decisions and the administration of benefits are subject to all terms, conditions, exclusions and

limitations of the coverage documents (e.g., evidence of coverage, certificate of coverage, policy, contract of insurance, etc.), as well as to state and federal requirements and applicable Health Plan-level administrative policies and procedures.

This clinical policy is effective as of the date determined by the Health Plan. The date of posting may not be the effective date of this clinical policy. This clinical policy may be subject to applicable legal and regulatory requirements relating to provider notification. If there is a discrepancy between the effective date of this clinical policy and any applicable legal or regulatory requirement, the requirements of law and regulation shall govern. The Health Plan retains the right to change, amend or withdraw this clinical policy, and additional clinical policies may be developed and adopted as needed, at any time.

This clinical policy does not constitute medical advice, medical treatment or medical care. It is not intended to dictate to providers how to practice medicine. Providers are expected to exercise professional medical judgment in providing the most appropriate care, and are solely responsible for the medical advice and treatment of members. This clinical policy is not intended to recommend treatment for members. Members should consult with their treating physician in connection with diagnosis and treatment decisions.

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Note:

For Medicaid members, when state Medicaid coverage provisions conflict with the coverage provisions in this clinical policy, state Medicaid coverage provisions take precedence. Please refer to the state Medicaid manual for any coverage provisions pertaining to this clinical policy.

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