

Clinical Policy: Benralizumab (Fasenra)

Reference Number: CP.PHAR.373

Effective Date: 06.01.18

Last Review Date: 11.25

Line of Business: Commercial*, Medicaid*

[Coding Implications](#)[Revision Log](#)

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

Description

Benralizumab (Fasenra[®]) is an interleukin-5 receptor alpha-directed cytolytic monoclonal antibody (IgG1 kappa).

**California Exchange Plans and NY CHIP Plans should not be approved using these criteria; for California Exchange Plans refer to the HIM.PA.SP70 Benralizumab (Fasenra) criteria; for NY CHIP Plans refer to the NY.HIM.PA.SP70 Benralizumab (Fasenra) criteria*

FDA Approved Indication(s)

Fasenra is indicated for the:

- Add-on maintenance treatment of patients with severe asthma aged 6 years and older, and with an eosinophilic phenotype
- Treatment of adult patients with eosinophilic granulomatosis with polyangiitis (EGPA)

Limitation(s) of use: Fasenra is not indicated for the relief of acute bronchospasm or status asthmaticus.

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 Fasenra is **medically necessary** when the following criteria are met:

I. Initial Approval Criteria**A. Severe Asthma*** (must meet all):

**Refer to HIM.PA.SP70 for California Exchange Plans and NY.HIM.PA.SP70 for NY CHIP Plans*

1. Diagnosis of asthma;
2. Member has an absolute blood eosinophil count ≥ 150 cells/mcL within the past 3 months;
3. Prescribed by or in consultation with a pulmonologist, immunologist, or allergist;
4. Age ≥ 6 years;
5. Member has experienced ≥ 2 exacerbations within the last 12 months, requiring one of the following (a or b), despite adherent use of controller therapy (i.e., medium- to high-dose inhaled corticosteroid [ICS] plus either a long-acting beta₂ agonist [LABA] or leukotriene modifier [LTRA] if LABA contraindication/intolerance):

- a. Oral/systemic corticosteroid treatment (or increase in dose if already on oral corticosteroid);
- b. Urgent care/emergency room (ER) visit or hospital admission;
6. Fasenra is prescribed concurrently with an ICS plus either a LABA or LTRA;
7. Fasenra is not prescribed concurrently with Cinqair[®], Nucala[®], Dupixent[®], Xolair[®], or Tezspire[®];
8. Dose does not exceed (a or b):
 - a. Age 6 to 11 years and weight < 35 kg: 10 mg every 4 weeks for the first 3 doses, then 10 mg every 8 weeks thereafter;
 - b. Age ≥ 6 years and weight ≥ 35 kg: 30 mg every 4 weeks for the first 3 doses, then 30 mg every 8 weeks thereafter.

Approval duration:

Medicaid – 12 months

Commercial – 6 months or member's renewal period, whichever is longer

B. Eosinophilic Granulomatosis with Polyangiitis (formerly Churg-Strauss) (must meet all):

1. Diagnosis of EGPA (formerly Churg-Strauss) with both of the following (a and b):
 - a. Active, non-severe disease;*
**Non-severe disease is defined as vasculitis without life- or organ-threatening manifestations. Examples of symptoms in patients with non-severe disease include rhinosinusitis, asthma, mild systemic symptoms, uncomplicated cutaneous disease, and mild inflammatory arthritis.*
 - b. Eosinophilia as evidenced by eosinophils > 1 x 10⁹/L and/or > 10% of leukocytes within the past 3 months;
2. Prescribed by or in consultation with a pulmonologist, rheumatologist, immunologist, or nephrologist;
3. Age ≥ 18 years;
4. Failure of a 4-week trial of a glucocorticoid (*see Appendix B*), unless contraindicated or clinically significant adverse events are experienced;
5. Fasenra is not prescribed concurrently with Cinqair, Nucala, Dupixent, Xolair, or Tezspire;
6. Dose does not exceed 30 mg every 4 weeks.

Approval duration:

Medicaid – 12 months

Commercial – 6 months or member's renewal period, whichever is longer

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

1. If this drug has recently (within the last 6 months) undergone a label change (e.g., newly approved indication, age expansion, new dosing regimen) that is not yet reflected in this policy, refer to one of the following policies (a or b):
 - a. For drugs on the formulary (commercial, health insurance marketplace) or PDL (Medicaid), the no coverage criteria policy for the relevant line of business: CP.CPA.190 for commercial and CP.PMN.255 for Medicaid; or
 - b. For drugs NOT on the formulary (commercial, health insurance marketplace) or PDL (Medicaid), the non-formulary policy for the relevant line of business: CP.CPA.190 for commercial and CP.PMN.16 for Medicaid; or

2. If the requested use (e.g., diagnosis, age, dosing regimen) is NOT specifically listed under section III (Diagnoses/Indications for which coverage is NOT authorized) AND criterion 1 above does not apply, refer to the off-label use policy for the relevant line of business: CP.CPA.09 for commercial and CP.PMN.53 for Medicaid.

II. Continued Therapy

A. Severe Asthma* (must meet all):

**Refer to HIM.PA.SP70 for California Exchange Plans and NY.HIM.PA.SP70 for NY CHIP Plans*

1. Member meets one of the following (a or b):
 - a. Currently receiving medication via Centene benefit or member has previously met initial approval criteria;
 - b. Member is currently receiving medication and is enrolled in a state and product with continuity of care regulations (*refer to state specific addendums for CC.PHARM.03A and CC.PHARM.03B*);
2. Demonstrated adherence to asthma controller therapy (an ICS plus either a LABA or LTRA) as evidenced by proportion of days covered (PDC) of 0.8 in the last 6 months (i.e., member has received asthma controller therapy for at least 5 of the last 6 months);
3. Member is responding positively to therapy (examples may include but are not limited to: reduction in exacerbations or corticosteroid dose, improvement in forced expiratory volume over one second since baseline, reduction in the use of rescue therapy);
4. Fasenra is not prescribed concurrently with Cinqair, Nucala, Dupixent, Xolair, or Tezspire;
5. If request is for a dose increase, new dose does not exceed (a or b):
 - a. Age 6 to 11 years and weight < 35 kg: 10 mg every 4 weeks for the first 3 doses, then 10 mg every 8 weeks thereafter;
 - b. Age ≥ 6 years and weight ≥ 35 kg: 30 mg every 4 weeks for the first 3 doses, then 30 mg every 8 weeks thereafter.

Approval duration:

Medicaid – 12 months

Commercial – 6 months or member's renewal period, whichever is longer

B. Eosinophilic Granulomatosis with Polyangiitis (formerly Churg-Strauss) (must meet all):

1. Member meets one of the following (a or b):
 - a. Currently receiving medication via Centene benefit or member has previously met initial approval criteria;
 - b. Member is currently receiving medication and is enrolled in a state and product with continuity of care regulations (*refer to state specific addendums for CC.PHARM.03A and CC.PHARM.03B*);
2. Member is responding positively to therapy (examples may include but are not limited to: reduction of relapses or reduction in glucocorticoid dose);
3. Fasenra is not prescribed concurrently with Cinqair, Nucala, Dupixent, Xolair, or Tezspire;

4. If request is for a dose increase, new dose does not exceed 30 mg every 4 weeks.

Approval duration:

Medicaid – 12 months

Commercial – 6 months or member's renewal period, whichever is longer

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

1. If this drug has recently (within the last 6 months) undergone a label change (e.g., newly approved indication, age expansion, new dosing regimen) that is not yet reflected in this policy, refer to one of the following policies (a or b):
 - a. For drugs on the formulary (commercial, health insurance marketplace) or PDL (Medicaid), the no coverage criteria policy for the relevant line of business: CP.CPA.190 for commercial and CP.PMN.255 for Medicaid; or
 - b. For drugs NOT on the formulary (commercial, health insurance marketplace) or PDL (Medicaid), the non-formulary policy for the relevant line of business: CP.CPA.190 for commercial and CP.PMN.16 for Medicaid; or
2. If the requested use (e.g., diagnosis, age, dosing regimen) is NOT specifically listed under section III (Diagnoses/Indications for which coverage is NOT authorized) AND criterion 1 above does not apply, refer to the off-label use policy for the relevant line of business: 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;
- B. Acute bronchospasm or status asthmaticus.

IV. Appendices/General Information

Appendix A: Abbreviation/Acronym Key

BEC: blood eosinophil count

EGPA: eosinophilic granulomatosis with polyangiitis

ER: emergency room

FDA: Food and Drug Administration

GINA: Global Initiative for Asthma

ICS: inhaled corticosteroid

LABA: long-acting beta2 agonist

LTRA: leukotriene modifier

PDC: proportion of days covered

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 and may require prior authorization.

Drug Name	Dosing Regimen	Dose Limit/ Maximum Dose
Asthma - ICS (medium – high dose)		
Qvar® (beclomethasone)	> 200 mcg/day 40 mcg, 80 mcg per actuation 1-4 actuations BID	4 actuations BID
budesonide (Pulmicort®)	> 400 mcg/day	2 actuations BID

Drug Name	Dosing Regimen	Dose Limit/ Maximum Dose
	90 mcg, 180 mcg per actuation 2-4 actuations BID	
Alvesco® (ciclesonide)	> 160 mcg/day 80 mcg, 160 mcg per actuation 1-2 actuations BID	2 actuations BID
fluticasone propionate (Flovent®)	> 250 mcg/day 44-250 mcg per actuation 2-4 actuations BID	2 actuations BID
Arnuity Ellipta® (fluticasone furoate)	200 mcg/day 100 mcg, 200 mcg per actuation 1 actuation QD	1 actuation QD
Asmanex® (mometasone)	> 200 mcg/day HFA: 100 mcg, 200 mcg per actuation Twisthaler: 110 mcg, 220 mcg per actuation 1-2 actuations QD to BID	2 inhalations BID
Asthma - LABA		
Serevent® (salmeterol)	50 mcg per dose 1 inhalation BID	1 inhalation BID
Asthma - Combination products (ICS + LABA)		
Dulera® (mometasone/ formoterol)	100/5 mcg, 200/5 mcg per actuation 2 actuations BID	4 actuations per day
Breo Ellipta® (fluticasone/vilanterol)	100/25 mcg, 200/25 mcg per actuation 1 actuation QD	1 actuation QD
fluticasone/salmeterol (Advair®)	Diskus: 100/50 mcg, 250/50 mcg, 500/50 mcg per actuation HFA: 45/21 mcg, 115/21 mcg, 230/21 mcg per actuation 1 actuation BID	1 actuation BID
fluticasone/salmeterol (Airduo RespiClick®)	55/13 mcg, 113/14 mcg, 232/14 mcg per actuation 1 actuation BID	1 actuation BID
budesonide/formoterol (Symbicort®)	80 mcg/4.5 mcg, 160 mcg/4.5 mcg per actuation 2 actuations BID	2 actuations BID
Asthma - LTRA		
montelukast (Singulair®)	4 to 10 mg PO QD	10 mg per day
zafirlukast (Accolate®)	10 to 20 mg PO BID	40 mg per day
zileuton ER (Zyflo® CR)	1,200 mg PO BID	2,400 mg per day
Zyflo® (zileuton)	600 mg PO QID	2,400 mg per day
Asthma - Oral corticosteroids		
dexamethasone (Decadron®)	0.75 to 9 mg/day PO in 2 to 4 divided doses	Varies

Drug Name	Dosing Regimen	Dose Limit/ Maximum Dose
methylprednisolone (Medrol®)	40 to 80 mg PO in 1 to 2 divided doses	Varies
prednisolone (Millipred®, Orapred ODT®)	40 to 80 mg PO in 1 to 2 divided doses	Varies
prednisone (Deltasone®)	40 to 80 mg PO in 1 to 2 divided doses	Varies
EGPA		
methylprednisolone (Medrol)	6.0 mg/day to 0.8 mg/kg/day	Varies
prednisone (Deltasone)	7.5 mg/day to 1 mg/kg/day	Varies
cyclophosphamide*	1-2 mg/kg/day PO or 0.5-1 g/m ² /month IV	See regimen
azathioprine*	2-3 mg/kg PO QD	See regimen
methotrexate*	15 mg/week PO	25 mg/week
mycophenolate mofetil*	1.5-3 g/day PO	3 g/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.

*Off-label

Appendix C: Contraindications/Boxed Warnings

- Contraindication(s): hypersensitivity
- Boxed warning(s): none reported

Appendix D: General Information

- Asthma:
 - The pivotal trials defined severe asthma as 2 or more exacerbations of asthma despite regular use of high-dose ICS plus an additional controller (e.g., LABA or LTRA) with or without oral corticosteroids. Although the CALIMA trial included patients receiving medium-dose ICS, Fasenera was not shown to have an effect on annual exacerbation rate, pre-bronchodilator forced expiratory volume in 1 second, or total asthma symptom score in those patients.
 - Clinically significant exacerbation was defined as a worsening of asthma (any new or increased symptoms or signs that were concerning) that led to one of the following: (1) use of systemic corticosteroids, (2) emergency department or visit to urgent care center, or (3) inpatient hospital stay.
 - Baseline blood eosinophil count (BEC) is a predictor of response to therapy. Although the SIROCCO and CALIMA trials were powered for efficacy analysis in patients with baseline BEC ≥ 300 cells/ μ L, a pooled analysis which stratified patients by baseline BEC (≥ 0 cells/ μ L, ≥ 150 cells/ μ L, ≥ 300 cells/ μ L, and ≥ 450 cells/ μ L) found Fasenera to have a statistically significant positive treatment effect on those with baseline BEC ≥ 150 cells/ μ L. In addition, the ZONDA trial found Fasenera to significantly reduce oral corticosteroid dose in patients with baseline BEC ≥ 150 cells/ μ L.
 - The Global Initiative for Asthma (GINA) guidelines recommend Fasenera be considered as adjunct therapy for patients 12 years of age and older with

- exacerbations or poor symptom control despite taking at least high dose ICS/LABA and who have eosinophilic biomarkers or need maintenance oral corticosteroids.
- Patients could potentially meet asthma criteria for both Xolair and Fasenra, though there is insufficient data to support the combination use of multiple asthma biologics. The combination has not been studied. Approximately 30% of patients in the Nucala MENSA study also were candidates for therapy with Xolair.
- Lab results for blood eosinophil counts can be converted into cells/mcL using the following unit conversion calculator: <https://www.fasenrahcp.com/m/fasenra-eosinophil-calculator.html>.
- PDC is a measure of adherence. PDC is calculated as the sum of days covered in a time frame divided by the number of days in the time frame. To achieve a PDC of 0.8, a member must have received their asthma controller therapy for 144 days out of the last 180 days, or approximately 5 months of the last 6 months.
- **EGPA:**
 - Standard of care for EGPA includes oral glucocorticoids. Induction therapy of prednisone 1 mg/kg/day is recommended for 2-3 weeks followed by gradual tapering to the minimal effective dose. Patients with stable doses of prednisone ≤ 7.5 mg/day are considered to be in remission, as defined by the European League Against Rheumatism (EULAR) and in the pivotal trial. The EGPA Consensus Task Force recommends that patients who are unable to taper prednisone to < 7.5 mg/day after 3-4 months of therapy should be considered for additional immunosuppressant therapy.

V. Dosage and Administration

Indication	Dosing Regimen	Maximum Dose
Severe asthma	<u>Adult and adolescents (12 years and older):</u> ● 30 mg SC every 4 weeks for the first 3 doses, followed by once every 8 weeks thereafter <u>Pediatric patients 6 - 11 years of age:</u> ● < 35 kg: 10 mg SC every 4 weeks for the first 3 doses, followed by once every 8 weeks thereafter ● ≥ 35 kg: 30 mg SC every 4 weeks for the first 3 doses, followed by once every 8 weeks thereafter	See regimen
EGPA	30 mg SC every 4 weeks	30 mg/4 weeks

VI. Product Availability

- Single-dose prefilled syringe with solution for injection: 10 mg/0.5 mL, 30 mg/mL
- Single-dose autoinjector Fasenra Pen with solution for injection: 30 mg/mL

VII. References

1. Fasenra Prescribing Information. Wilmington, DE: AstraZeneca Pharmaceuticals LP; September 2024. Available at: www.fasenra.com. Accessed October 24, 2024.

2. Clinical Pharmacology [database online]. Philadelphia, PA: Elsevier. Updated periodically. Available at: <http://www.clinicalkey.com/pharmacology>. Accessed November 14, 2024.

Asthma

3. National Asthma Education and Prevention Program: Expert panel report III: Guidelines for the diagnosis and management of asthma. Bethesda, MD: National Heart, Lung, and Blood Institute, 2007. (NIH publication no. 08-4051). Available at <http://www.nhlbi.nih.gov/health-pro/guidelines/current/asthma-guidelines>. Accessed November 14, 2024.
4. Cloutier MM, Dixon AE, Krishnan JA, et al. Managing asthma in adolescents and adults 2020: asthma guideline update from the National Asthma Education and Prevention Program. JAMA. 2020; 324: 2301-2317.
5. Global Initiative for Asthma. Global strategy for asthma management and prevention (2024 update). Available from: www.ginasthma.org. Accessed November 14, 2024.
6. Global Initiative for Asthma. Difficult-to-treat and severe asthma in adolescent and adult patients – diagnosis and management, v5.0 November 2024. Available at: www.ginasthma.org. Accessed November 14, 2024.

EGPA

7. Wechsler ME, Nair P, Terrier B, et al. Benralizumab versus mepolizumab for eosinophilic granulomatosis with polyangiitis. N Engl J Med. 2024; 390: 911-921.
8. Chung SA, Langford CA, Maz M, et al. 2021 American College of Rheumatology/Vasculitis Foundation guideline for the management of antineutrophil cytoplasmic antibody-associated vasculitis. Arthritis Care & Research. 2021; 73(8): 1088-1105.
9. Grayson PC, Ponte C, Suppiah R, et al. 2022 American College of Rheumatology/European Alliance of Associations for Rheumatology classification criteria for eosinophilic granulomatosis with polyangiitis. Annals of the Rheumatic Diseases. 2022; 81: 309-314.

Coding Implications

Codes referenced in this clinical policy are for informational purposes only. Inclusion or exclusion of any codes does not guarantee coverage. Providers should reference the most up-to-date sources of professional coding guidance prior to the submission of claims for reimbursement of covered services.

HCPCS Codes	Description
J0517	Injection, benralizumab, 1 mg

Reviews, Revisions, and Approvals	Date	P&T Approval Date
1Q 2021 annual review: no significant changes; references to HIM.PHAR.21 revised to HIM.PA.154; HCPCS code updated; references reviewed and updated.	10.26.20	02.21
1Q 2022 annual review: for continuation criteria, defined adherence as PDC of 0.8; references reviewed and updated.	09.22.21	02.22
Template changes applied to other diagnoses/indications and continued therapy section.	09.22.22	

Reviews, Revisions, and Approvals	Date	P&T Approval Date
1Q 2023 annual review: no significant changes; added Tezspire as another agent with which Fasenra should not be used concurrently; references reviewed and updated.	10.31.22	02.23
1Q 2024 annual review: no significant changes; references reviewed and updated.	11.05.23	02.24
RT4: expanded age to 6+ years old and added new 10 mg prefilled syringe formulation; references reviewed and updated.	04.26.24	
Per SDC, removed HIM line of business as separate criteria is required; added reference to “Refer to HIM.PA.SP70 for California Exchange Plans”; for initial approval criteria, added allowance for ER visit. RT4: added criteria for newly approved indication of EGPA.	10.23.24	11.24
1Q 2025 annual review: no significant changes; references reviewed and updated. Per December SDC: added statement disclaimer that NY CHIP Plans should not be approved using these criteria and should use NY.HIM.PA.SP70 criteria; for asthma, removed intubation option for alignment purposes as a hospital admission would encompass intubation.	12.02.24	02.25
For both indications, revised initial approval duration for Medicaid from 6 to 12 months and for Commercial to include “or to the member’s renewal date, whichever is longer” for this maintenance medication for a chronic condition.	10.21.25	11.25

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.

Providers referred to in this clinical policy are independent contractors who exercise independent judgment and over whom the Health Plan has no control or right of control. Providers are not agents or employees of the Health Plan.

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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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