

Xolair® (omalizumab) (Subcutaneous)

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I. Length of Authorization

Coverage will be provided for six months and may be renewed (Management of Immune Checkpoint Inhibitor-Related Toxicity may NOT be renewed)

II. Dosing Limits

A. Quantity Limit (max daily dose) [Pharmacy Benefit]:

- Xolair 75 mg single-dose prefilled syringe: 1 syringe every 14 days
- Xolair 150 mg single-dose prefilled syringe: 2 syringes every 14 days
- Xolair 150mg powder for injection: 3 vials every 14 days

B. Max Units (per dose and over time) [Medical Benefit]:

Allergic Asthma

- 90 billable units every 14 days

All other indications

- 60 billable units every 28 days

III. Initial Approval Criteria

Coverage is provided in the following conditions:

- Must not be used in combination with another monoclonal antibody (e.g., benralizumab, mepolizumab, reslizumab, etc.); **AND**
- Patient must be at least 18 years of age (unless otherwise specified); **AND**

Moderate-to-severe persistent allergic asthma †

- Patient must be at least 6 years of age; **AND**
- Patient has a positive skin test or in vitro reactivity to a perennial aeroallergen; **AND**
- Patient must weigh between 20 kg (44 lbs.) and 150 kg (330 lbs.); **AND**
- Patient has a serum total IgE level, measured before the start of treatment, of either:

- ≥ 30 IU/mL and ≤ 700 IU/mL in patients age ≥ 12 years; **OR**
- ≥ 30 IU/mL and ≤ 1300 IU/mL in patients age 6 to <12 years; **AND**
- Patient has documented ongoing symptoms of moderate-to-severe asthma* with a minimum (3) month trial on previous combination therapy including medium- or high-dose inhaled corticosteroids **PLUS** another controller medication (e.g., long-acting beta-2 agonist, leukotriene receptor antagonist, theophylline, etc.)

Chronic idiopathic urticaria (CIU) †

- Patient must be at least 12 years of age; **AND**
- The underlying cause of the patient's condition is NOT considered to be any other allergic condition(s) or other form(s) of urticaria; **AND**
- Patient is avoiding triggers (e.g., NSAIDs, etc.); **AND**
- Documented baseline score from an objective clinical evaluation tool, such as: urticaria activity score (UAS7), angioedema activity score (AAS), Dermatology Life Quality Index (DLQI), Angioedema Quality of Life (AE-QoL), or Chronic Urticaria Quality of Life Questionnaire (CU-Q₂₀L); **AND**
- Patient had an inadequate response to a one or more month trial on previous therapy with scheduled dosing of a second-generation H1-antihistamine product**; **AND**
- Patient had an inadequate response to a one or more month trial on previous therapy with scheduled dosing of at least one of the following:
 - Updosing/dose advancement (up to 4-fold) of a second generation H1-antihistamine**
 - Add-on therapy with a leukotriene antagonist (e.g., montelukast, zafirlukast, etc.)
 - Add-on therapy with another H1-antihistamine**
 - Add-on therapy with a H2-antagonist (e.g. ranitidine, etc.)
 - Add-on therapy with cyclosporine

Note: renewal will require submission of a current (within 30 days) score from an objective clinical evaluation tool (i.e., UAS7, AAS, DLQI, AE-QoL or CU-Q₂₀L).

Management of Immune Checkpoint Inhibitor-Related Toxicity ‡

- Patient has been receiving therapy with an immune checkpoint inhibitor (e.g. nivolumab, pembrolizumab, atezolizumab, avelumab, durvalumab, cemiplimab, ipilimumab, etc.); **AND**
- Patient has refractory and severe (i.e., grade 3: intense or widespread, constant, limiting self-care activities of daily living or sleep) pruritis; **AND**
- Patient has an increased serum IgE level above the upper limit of normal of the laboratory reference value

Systemic Mastocytosis ‡

- Used for the prevention of one of the following:

- Chronic mast-cell-mediator-related cardiovascular (e.g., pre-syncope, tachycardia, etc.) or pulmonary (e.g., wheezing, throat-swelling, etc.) symptoms insufficiently controlled by conventional therapy (e.g., H1 or H2 blockers or corticosteroids); **OR**
- Unprovoked anaphylaxis; **OR**
- Hymenoptera or food-induced anaphylaxis in patients with a negative test for specific IgE antibodies or a negative skin test; **OR**
- Used to improve tolerance while on immunotherapy (i.e., venom immunotherapy [VIT])

***Components of severity for classifying asthma as moderate may include any of the following (not all inclusive):**

- Daily symptoms
- Nighttime awakenings > 1x/week but not nightly
- SABA use for symptom control occurs daily
- Some limitation to normal activities
- Lung function (percent predicted FEV₁) >60%, but <80%
- Exacerbations requiring oral systemic corticosteroids are generally more frequent and intense relative to mild asthma

***Components of severity for classifying asthma as severe may include any of the following (not all inclusive):**

- Symptoms throughout the day
- Nighttime awakenings, often 7x/week
- SABA use for symptom control occurs several times daily
- Extremely limited in normal activities
- Lung function (percent predicted FEV₁) <60%
- Exacerbations requiring oral systemic corticosteroids are generally more frequent and intense relative to moderate asthma

****H1 Antihistamine Products (not all inclusive)**

- fexofenadine
- loratadine
- desloratadine
- cetirizine
- levocetirizine
- clemastine
- diphenhydramine
- chlorpheniramine
- hydroxyzine
- cyproheptadine
- brompheniramine
- triprolidine
- dexchlorpheniramine
- carbinoxamine

† FDA-approved indication(s); ‡ Compendia recommended indication(s)

IV. Renewal Criteria

- Patient continues to meet the criteria identified in section III; **AND**
- Absence of unacceptable toxicity from the drug. Examples of unacceptable toxicity include the following: symptoms of anaphylaxis (bronchospasm, hypotension, syncope, urticaria,

and/or angioedema), malignancy, symptoms similar to serum sickness (fever, arthralgia, and rash); parasitic (helminth) infection, eosinophilic conditions (e.g. vasculitic rash, worsening pulmonary symptoms, cardiac complications, and/or neuropathy, especially upon reduction of oral corticosteroids), etc.; **AND**

Moderate-to-severe persistent allergic asthma

- Patient must weigh between 20 kg (44 lbs.) and 150 kg (330 lbs.); **AND**
- Treatment has resulted in clinical improvement as documented by one or more of the following:
 - Decreased utilization of rescue medications; **OR**
 - Decreased frequency of exacerbations (defined as worsening of asthma that requires increase in inhaled corticosteroid dose or treatment with systemic corticosteroids); **OR**
 - Improvement in lung function (increase in percent predicted FEV₁ or PEF) from pre-treatment baseline; **OR**
 - Reduction in reported symptoms (e.g., decrease in asthma symptom score), as evidenced by decreases in frequency or magnitude of one or more of the following symptoms:
 - Asthma attacks
 - Chest tightness or heaviness
 - Coughing or clearing throat
 - Difficulty taking deep breath or difficulty breathing out
 - Shortness of breath
 - Sleep disturbance, night wakening, or symptoms upon awakening
 - Tiredness
 - Wheezing/heavy breathing/fighting for air; **AND**
- Patient is periodically checked to reassess the need for continued therapy based upon the patient's disease severity and level of asthma control

Chronic idiopathic urticaria (CIU)

- Treatment with Xolair (omalizumab) has resulted in clinical improvement as documented by improvement from baseline using objective clinical evaluation tools such as the urticaria activity score (UAS7), angioedema activity score (AAS), Dermatology Life Quality Index (DLQI), Angioedema Quality of Life (AE-QoL), or Chronic Urticaria Quality of Life Questionnaire(CU-Q₂₀L); **AND**
- Submitted current UAS7, AAS, DLQI, AE-QoL, or Cu-Q₂₀L was recorded within the past 30 days.

Management of Immune Checkpoint Inhibitor related Toxicity

- May not be renewed

Systemic Mastocytosis

- Disease response as indicated by improvement in signs and symptoms compared to baseline or a decreased frequency of exacerbations

V. Dosage/Administration

Indication	Dose
Allergic Asthma	75 to 375 mg administered subcutaneously by a health care provider every 2 or 4 weeks. Determine dose (mg) and dosing frequency by serum total IgE level (IU/mL), measured before the start of treatment, and body weight (kg). See tables below.
Chronic idiopathic urticaria	150 or 300 mg administered subcutaneously by a health care provider every 4 weeks. Dosing is not dependent on serum IgE (free or total) level or body weight.
All other indications	150 or 300 mg administered subcutaneously by a health care provider every 4 weeks. Dosing is not dependent on serum IgE (free or total) level or body weight.

Omaliuzumab Doses Administered Every 4 Weeks (mg) in patients ≥ 12 years				
Pre-treatment serum IgE (IU/mL)	Body weight (kg)			
	30 to 60	> 60 to 70	> 70 to 90	> 90 to 150
≥ 30 to 100	150	150	150	300
> 100 to 200	300	300	300	See the following table.
> 200 to 300	300	See the following table.	See the following table.	See the following table.

Omaliuzumab Doses Administered Every 2 Weeks (mg) in patients ≥ 12 years				
Pre-treatment serum IgE (IU/mL)	Body weight (kg)			
	30 to 60	> 60 to 70	> 70 to 90	> 90 to 150
> 100 to 200	See previous table.	See previous table.	See previous table.	225
> 200 to 300	See previous table.	225	225	300
> 300 to 400	225	225	300	Do not dose.
> 400 to 500	300	300	375	Do not dose.
> 500 to 600	300	375	Do not dose.	Do not dose.
> 600 to 700	375	Do not dose.	Do not dose.	Do not dose

Omaliuzumab Doses Administered Every 2 or 4 Weeks (mg) for Pediatric Patients with Asthma Who Begin Xolair Between the Ages of 6 to <12 Years											
Pre-treatment IgE (IU/mL)	Dosing Freq. (weeks)	Body Weight (kg)									
		20-25	>25-30	>30-40	>40-50	>50-60	>60-70	>70-80	>80-90	>90-125	>125-150

30-100	4	75	75	75	150	150	150	150	150	300	300	
>100-200		150	150	150	300	300	300	300	300	225	300	
>200-300		150	150	225	300	300	225	225	225	300	375	
>300-400		225	225	300	225	225	225	300	300	Do Not Dose		
>400-500		225	300	225	225	300	300	375	375			
>500-600		300	300	225	300	300	375					
>600-700		300	225	225	300	375						
>700-900	2	225	225	300	375							
>900-1100		225	300	375								
>1100-1200		300	300									
>1200-1300		300	375									

VI. Billing Code/Availability Information

Jcode:

- J2357 – Injection, omalizumab, 5 mg; 1 billable unit = 5 mg

NDC:

- Xolair 75 mg single-dose prefilled syringe: 50242-0214-xx
- Xolair 150 mg single-dose prefilled syringe: 50242-0215-xx
- Xolair 150 mg single-use vial powder for injection: 50242-0040-xx

VII. References

1. Xolair [package insert]. South San Francisco, CA; Genentech, Inc.; May 2019. Accessed September 2019.
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3. Global Initiative for Asthma (GINA). Global Strategy for Asthma Management and Prevention. 2018 Update. Available from: <http://www.ginasthma.org>. Accessed April 2018.
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LEN taskforce position paper. *Allergy*. 2011 Jul;66(7):840-4. doi: 10.1111/j.1398-9995.2011.02580.x. Epub 2011 Mar 9.

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6. Maurer M, Rosén K, Hsieh HJ, et al. Omalizumab for the treatment of chronic idiopathic or spontaneous urticaria. *N Engl J Med*. 2013 Mar 7;368(10):924-35. doi: 10.1056/NEJMoa1215372. Epub 2013 Feb 24.
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8. Bernstein JA, Lang DM, Khan DA, et al. The diagnosis and management of acute and chronic urticaria: 2014 update. *J Allergy Clin Immunol*. 2014 May;133(5):1270-7.
9. First Coast Service Options, Inc. Local Coverage Determination (LCD): Omalizumab (Xolair) (L33924). Centers for Medicare & Medicare Services. Updated on 02/01/2019 with effective dates 1/22/2019. Accessed September 2019.
10. National Government Services, Inc. Local Coverage Article: Omalizumab (e.g., Xolair) – Related to LCD L33394 (A52448). Centers for Medicare & Medicare Services. Updated on 12/24/2015 with effective dates 10/01/2015. Accessed September 2019.

Appendix 1 – Covered Diagnosis Codes

ICD-10	ICD-10 Description
C96.20	Malignant mast cell neoplasm, unspecified
C96.21	Aggressive systemic mastocytosis
C96.22	Mast cell sarcoma
C96.29	Other malignant mast cell neoplasm
D47.02	Systemic mastocytosis
J45.40	Moderate persistent asthma, uncomplicated
J45.41	Moderate persistent asthma with (acute) exacerbation
J45.42	Moderate persistent asthma with status asthmaticus
J45.50	Severe persistent asthma, uncomplicated
J45.51	Severe persistent asthma with (acute) exacerbation
J45.52	Severe persistent asthma with status asthmaticus
L29.8	Other pruritus
L29.9	Pruritus, unspecified
L50.1	Idiopathic urticaria

Appendix 2 – Centers for Medicare and Medicaid Services (CMS)

Medicare coverage for outpatient (Part B) drugs is outlined in the Medicare Benefit Policy Manual (Pub. 100-2), Chapter 15, §50 Drugs and Biologicals. In addition, National Coverage Determination (NCD) and Local Coverage Determinations (LCDs) may exist and compliance with these policies is required where applicable. They can be found at: <http://www.cms.gov/medicare-coverage-database/search/advanced-search.aspx>. Additional indications may be covered at the discretion of the health plan.

Jurisdiction(s): N (9)	NCD/LCD Document (s): L33924
https://www.cms.gov/medicare-coverage-database/search/lcd-date-search.aspx?DocID=L33924&bc=gAAAAAAAAAAAAAAAAA==	
Jurisdiction(s): 6, K	NCD/LCD Document (s): A52448
https://www.cms.gov/medicare-coverage-database/search/article-date-search.aspx?DocID=A52448&bc=gAAAAAAAAAAAAAAAAA==	

Medicare Part B Administrative Contractor (MAC) Jurisdictions		
Jurisdiction	Applicable State/US Territory	Contractor
E (1)	CA, HI, NV, AS, GU, CNMI	Noridian Healthcare Solutions, LLC
F (2 & 3)	AK, WA, OR, ID, ND, SD, MT, WY, UT, AZ	Noridian Healthcare Solutions, LLC
5	KS, NE, IA, MO	Wisconsin Physicians Service Insurance Corporation (WPS)
6	MN, WI, IL	National Government Services, Inc. (NGS)
H (4 & 7)	LA, AR, MS, TX, OK, CO, NM	Novitas Solutions, Inc.
8	MI, IN	Wisconsin Physicians Service Insurance Corporation (WPS)
N (9)	FL, PR, VI	First Coast Service Options, Inc.
J (10)	TN, GA, AL	Palmetto GBA, LLC
M (11)	NC, SC, WV, VA (excluding below)	Palmetto GBA, LLC
L (12)	DE, MD, PA, NJ, DC (includes Arlington & Fairfax counties and the city of Alexandria in VA)	Novitas Solutions, Inc.
K (13 & 14)	NY, CT, MA, RI, VT, ME, NH	National Government Services, Inc. (NGS)
15	KY, OH	CGS Administrators, LLC