

# SCIG (immune globulin SQ): Hizentra<sup>®</sup>, Gammagard Liquid<sup>®</sup>, Gamunex<sup>®</sup>-C, Gammaked<sup>®</sup>, Hyqvia<sup>®</sup>, Cuvitru<sup>®</sup>, Cutaquig<sup>®</sup>, Xembify<sup>®</sup> (Subcutaneous)

Document Number: MODA-0059

Last Review Date: 07/01/2022

Date of Origin: 7/20/2010

Dates Reviewed: 09/2010, 12/2010, 03/2011, 06/2011, 09/2011, 12/2011, 03/2012, 06/2012, 09/2012, 12/2012, 03/2013, 06/2013, 09/2013, 12/2013, 03/2014, 09/2014, 12/2014, 03/2015, 06/2015, 09/2015, 12/2015, 03/2016, 06/2016, 09/2016, 12/2016, 03/2017, 06/2017, 09/2017, 12/2017, 03/2018, 04/2018, 06/2018, 10/2018, 01/2019, 08/2019, 10/2019, 10/2020, 10/2021, 12/2021, 07/2022

## I. Length of Authorization

Initial coverage will be provided for 6 months and may be renewed annually thereafter.

## II. Dosing Limits

### A. Quantity Limit (max daily dose) [NDC Unit]:

Drug Name	Dose/ week	Dose/28 days
Hizentra	46 g	184 g
Gamunex-C & Gammaked	24 g	96 g
Gammagard liquid	24 g	96 g
HyQvia	17.5 g	69 g
Cuvitru	23 g	92 g
Cutaquig	24 g	96 g
Xembify	24 g	96 g

### B. Max Units (per dose and over time) [HCPCS Unit]:

Drug Name	Billable units/28 days
Hizentra	960 (PID)
	1840 (CIDP)
Gamunex-C & Gammaked	192
Gammagard liquid	192
HyQvia	690
Cuvitru	920
Cutaquig	960
Xembify	960

### III. Initial Approval Criteria <sup>1-8, 15,18</sup>

Site of care specialty infusion program requirements are met (refer to [Moda Site of Care Policy](#)).

Coverage is provided in the following conditions:

- Baseline values for BUN and serum creatinine obtained within 30 days of request; **AND**

#### **Primary immunodeficiency (PID)/Wiskott -Aldrich syndrome †**

Such as: x-linked agammaglobulinemia, common variable immunodeficiency, transient hypogammaglobulinemia of infancy, IgG subclass deficiency with or without IgA deficiency, antibody deficiency with near normal immunoglobulin levels) and combined deficiencies (severe combined immunodeficiencies, ataxia-telangiectasia, x-linked lymphoproliferative syndrome) ***[list not all inclusive]***

- Patient is  $\geq 2$  years old [HyQvia ONLY: patient must be  $\geq 18$  years old]; **AND**
- Patient's IgG level is  $<200$  mg/dL **OR both** of the following:
  - Patient has a history of multiple hard to treat infections as indicated by at least **one** of the following:
    - Four or more ear infections within 1 year
    - Two or more serious sinus infections within 1 year
    - Two or more months of antibiotics with little effect
    - Two or more pneumonias within 1 year
    - Recurrent or deep skin abscesses
    - Need for intravenous antibiotics to clear infections
    - Two or more deep-seated infections including septicemia; **AND**
  - The patient has a deficiency in producing antibodies in response to vaccination; **AND**
    - Titers were drawn before challenging with vaccination; **AND**
    - Titers were drawn between 4 and 8 weeks of vaccination

#### **Chronic Inflammatory Demyelinating Polyneuropathy (CIDP) [Hizentra ONLY] † Φ**

- Patient must be  $\geq 18$  years old; **AND**
- Physician has assessed baseline disease severity utilizing an objective measure/tool (e.g., INCAT, Medical Research Council (MRC) muscle strength, 6-MWT, Rankin, Modified Rankin, etc.); **AND**
  - Used as initial maintenance therapy for prevention of disease relapses after treatment and stabilization with intravenous immunoglobulin (IVIG)§; **OR**
  - Used for re-initiation of maintenance therapy after experiencing a relapse and requiring re-induction therapy with IVIG (see Section IV for criteria)

#### **Acquired Immune Deficiency secondary to Chronic Lymphocytic Leukemia ‡ <sup>31,32</sup>**

- Patient's IgG level is <200 mg/dL **OR both** of the following:
  - Patient has a history of multiple hard to treat infections as indicated by at least **one** of the following:
    - Four or more ear infections within 1 year
    - Two or more serious sinus infections within 1 year
    - Two or more months of antibiotics with little effect
    - Two or more pneumonias within 1 year
    - Recurrent or deep skin abscesses
    - Need for intravenous antibiotics to clear infections
    - Two or more deep-seated infections including septicemia; **AND**
  - The patient has a deficiency in producing antibodies in response to vaccination; **AND**
    - Titers were drawn before challenging with vaccination; **AND**
    - Titers were drawn between 4 and 8 weeks of vaccination

Note: other secondary immunodeficiencies resulting in hypogammaglobulinemia and/or B-cell aplasia will be evaluated on a case-by-case basis

*§ Refer to the Immune Globulins medical necessity criteria (Document Number: IC-0071) for the relevant intravenous criteria requirements*

† FDA Approved Indication(s); ‡ Compendia Recommended Indication(s); ◊ Orphan Drug

#### IV. **Renewal Criteria** <sup>1-8, 15,18</sup>

Coverage can be renewed for 1 year based upon the following criteria:

- Patient continues to meet the universal and other indication-specific relevant criteria identified in section III; **AND**
- Absence of unacceptable toxicity from the drug. Examples of unacceptable toxicity include: severe hypersensitivity/anaphylaxis, thrombosis, aseptic meningitis syndrome, hemolytic anemia, hyperproteinemia, acute lung injury, etc.; **AND**
- BUN and serum creatinine obtained within the last 6 months and the concentration and rate of infusion have been adjusted accordingly; **AND**

##### **Primary immunodeficiency (PID)/Wiskott -Aldrich syndrome**

- Disease response as evidenced by one or more of the following:
  - Decrease in the frequency of infection
  - Decrease in the severity of infection

##### **Chronic Inflammatory Demyelinating Polyneuropathy (CIDP) [Hizentra ONLY]**

- Renewals will be authorized for patients that have demonstrated a beneficial clinical response to maintenance therapy, without relapses, based on an objective clinical measuring tool (e.g., INCAT, Medical Research Council (MRC) muscle strength, 6-MWT, Rankin, Modified Rankin, etc.); **OR**
- Patient is re-initiating maintenance therapy after experiencing a relapse while on Hizentra; **AND**
  - Patient improved and stabilized on IVIG treatment: **AND**

- Patient was NOT receiving maximum dosing of Hizentra prior to relapse

**Acquired Immune Deficiency secondary to Chronic Lymphocytic Leukemia** <sup>31,32</sup>

- Disease response as evidenced by one or more of the following:
  - Decrease in the frequency of infection
  - Decrease in the severity of infection; **AND**
- Patient is at a decreased risk of infection as a result of treatment necessitating continued therapy

**V. Dosage/Administration**

Dosing should be calculated using adjusted body weight if one or more of the following criteria are met:

- Patient’s body mass index (BMI) is 30 kg/m<sup>2</sup> or more; **OR**
- Patient’s actual body weight is 20% higher than his or her ideal body weight (IBW)

Use the following dosing formulas to calculate the adjusted body weight (round dose to nearest 5 gram increment in adult patients)
<b>Dosing formulas</b>
BMI = 703 x (weight in pounds/height in inches <sup>2</sup> )
IBW(kg) for males = 50 + [2.3 (height in inches – 60)]
IBW(kg) for females = 45.5 + [2.3 x (height in inches – 60)]
Adjusted body weight = IBW + 0.5 (actual body weight – IBW)

*This information is not meant to replace clinical decision making when initiating or modifying medication therapy and should only be used as a guide. Patient-specific variables should be taken into account.*

Indication	Dose
Chronic Inflammatory Demyelinating Polyneuropathy	<p><u>Hizentra ONLY:</u></p> <ul style="list-style-type: none"> <li>▪ Initiate therapy 1 week after the last IVIG dose</li> <li>▪ The recommended subcutaneous dose is 0.2 g/kg (1 mL/kg) body weight per week, administered in 1 or 2 sessions over 1 or 2 consecutive days.</li> <li>▪ If CIDP symptoms worsen, consider increasing the dose to 0.4 g/kg (2 mL/kg) body weight per week, administered in 2 sessions over 1 or 2 consecutive days.</li> <li>▪ If CIDP symptoms worsen on the 0.4 g/kg body weight per week dose, consider re-initiating therapy with an IVIG while discontinuing Hizentra.</li> </ul>

Indication	Dose
Primary immune deficiency including Wiskott-Aldrich Syndrome AND Acquired Immune Deficiency secondary to Chronic Lymphocytic Leukemia	<u>Hizentra:</u> <ul style="list-style-type: none"> <li>▪ Switching from IVIG               <ul style="list-style-type: none"> <li>○ Initiate therapy 1 week after the last IVIG dose</li> <li>○ Weekly dose: <math>1.37 \times (\text{previous IVIG dose (g)} / \text{number of weeks between IVIG doses})</math></li> <li>○ May be administered from daily up to every two weeks (biweekly)</li> <li>○ Biweekly dose: twice the weekly dose (using calculation above)</li> <li>○ Frequent dosing (2-7 times per week): divide the calculated weekly dose by the desired number of times per week</li> </ul> </li> <li>▪ Switching from SCIG               <ul style="list-style-type: none"> <li>○ Initiate therapy 1 week after the last SCIG dose</li> <li>○ Weekly dose (in grams) should be same as the weekly dose of prior SCIG treatment (in grams)</li> <li>○ Biweekly dose: multiply the calculated weekly dose by 2</li> <li>○ Frequent dosing (2-7 times per week): divide the calculated weekly dose by the desired number of times per week</li> </ul> </li> </ul>
	<u>Gamunex-C/Gammaked/Gammagard Liquid:</u> <ul style="list-style-type: none"> <li>▪ Initiate therapy 1 week after the last IVIG dose</li> <li>▪ Weekly dose: <math>1.37 \times (\text{previous IVIG dose (g)} / \text{number of weeks between IVIG doses})</math></li> </ul>
	<u>HyQvia:</u> <ul style="list-style-type: none"> <li>▪ Naïve to IgG or switching from SCIG: 300 to 600 mg/kg at 3 to 4 week intervals after initial ramp-up*</li> <li>▪ Switching from IGIV: use the same dose and frequency as the previous IV treatment after initial ramp-up*</li> </ul> <p><b>NOTE:</b> For patients previously on another IgG treatment, initiate therapy 1 week after the last infusion of IVIG or SCIG</p>
	<u>Xembify:</u> <ul style="list-style-type: none"> <li>▪ Switching from IVIG               <ul style="list-style-type: none"> <li>○ Start treatment one week after the last IVIG infusion.</li> <li>○ Weekly dose: <math>1.37 \times (\text{previous monthly (or every 3- week) IVIG dose in grams} / \text{number of weeks between IVIG doses})</math></li> </ul> </li> <li>▪ Switching from SCIG               <ul style="list-style-type: none"> <li>○ Weekly dose (in grams) should be same as the weekly dose of prior SCIG treatment (in grams)</li> </ul> </li> </ul>
	<u>Cuvitru:</u> <ul style="list-style-type: none"> <li>▪ Switching from IVIG or HyQvia               <ul style="list-style-type: none"> <li>○ Initiate therapy 1 week after the last IVIG or Hyqvia dose</li> <li>○ Weekly dose: <math>1.30 \times (\text{previous IVIG or HyQvia dose (g)} / \text{number of weeks between IVIG or HyQvia doses})</math></li> <li>○ May be administered from daily up to every two weeks (biweekly)</li> </ul> </li> </ul>

Indication	Dose
	<ul style="list-style-type: none"> <li>○ Biweekly dose: twice the weekly dose (using calculation above)</li> <li>○ Frequent dosing (2-7 times per week): divide the calculated weekly dose by the desired number of times per week</li> <li>▪ Switching from SCIG               <ul style="list-style-type: none"> <li>○ Weekly dose (in grams) should be same as the weekly dose of prior SCIG treatment (in grams)</li> <li>○ May be administered from daily up to every two weeks (biweekly)</li> <li>○ Biweekly dose: multiply the calculated weekly dose by 2</li> <li>○ Frequent dosing (2-7 times per week): divide the calculated weekly dose by the desired number of times per week</li> </ul> </li> </ul>
	<p data-bbox="384 590 505 617"><u>Cutaquig:</u></p> <p data-bbox="384 632 1479 699"><b>NOTE:</b> Start treatment one week after the last IVIG or SCIG infusion. Ensure that patients have received IVIG or SCIG treatment at regular intervals for at least 3 months</p> <ul style="list-style-type: none"> <li>▪ Switching from IVIG               <ul style="list-style-type: none"> <li>○ Weekly dose: <math>1.30 \times (\text{previous IVIG dose (g)} / \text{number of weeks between IVIG doses})</math></li> <li>○ May be administered from daily up to every two weeks (biweekly)</li> <li>○ Biweekly dose: multiply the calculated weekly dose by 2</li> <li>○ Frequent dosing (2-7 times per week): divide the calculated weekly dose by the desired number of times per week</li> </ul> </li> <li>▪ Switching from SCIG               <ul style="list-style-type: none"> <li>○ Weekly dose (in grams) should be same as the weekly dose of prior SCIG treatment (in grams)</li> <li>○ May be administered from daily up to every two weeks (biweekly)</li> <li>○ Biweekly dose: multiply the calculated weekly dose by 2</li> <li>○ Frequent dosing (2-7 times per week): divide the calculated weekly dose by the desired number of times per week</li> </ul> </li> </ul>

*Dosing for immunoglobulin products is highly variable depending on numerous patient specific factors, indication(s), and the specific product selected. For specific dosing regimens refer to current prescribing literature.*

\*HyQvia initial treatment interval/dosage ramp-up schedule

Week	Infusion Number	3-week treatment interval	4-week treatment interval
1	1 <sup>st</sup> infusion	Dose in Grams X 0.33	Dose in Grams X 0.25
2	2 <sup>nd</sup> infusion	Dose in Grams X 0.67	Dose in Grams X 0.50
4	3 <sup>rd</sup> infusion	Total Dose in Grams	Dose in Grams X 0.75
7	4 <sup>th</sup> infusion	N/A	Total Dose in Grams

## VI. Billing Code/Availability Information

HCP Code & NDC(s):

Drug Name*	Manufacturer	HCP Code	1 Billable unit	NDC	IgG (grams) per SDV	Volume (mL)
Hizentra 20% (Vials)	CSL Behring AG	J1559 – Injection, immune globulin (Hizentra), 100 mg	100 mg	44206-0451-01	1	5
				44206-0452-02	2	10
				44206-0454-04	4	20
				44206-0455-10	10	50
Hizentra 20% (Prefilled Syringes)	CSL Behring AG	J1559 – Injection, immune globulin (Hizentra), 100 mg	100 mg	44206-0456-21	1	5
				44206-0457-22	2	10
				44206-0458-24	4	20
Gammaked 10%	Grifols Therapeutics	J1561 – Injection, immune globulin, (Gamunex-C/ Gammaked), non-lyophilized (e.g., liquid), 500 mg	500 mg	76125-0900-01	1	10
				76125-0900-25	2.5	25
				76125-0900-50	5	50
				76125-0900-10	10	100
				76125-0900-20	20	200
Gamunex-C 10%	Grifols Therapeutics	J1561 – Injection, immune globulin, (Gamunex-C/Gammaked), non-lyophilized (e.g., liquid), 500 mg	500 mg	13533-0800-12	1	10
				13533-0800-15	2.5	25
				13533-0800-20	5	50
				13533-0800-71	10	100
				13533-0800-24	20	200
				13533-0800-40	40	400
Gammagard Liquid 10%	Baxalta US Inc.	J1569 – Injection, immune globulin, (Gammagard liquid), non-lyophilized, (e.g., liquid), 500 mg	500 mg	00944-2700-02	1	10
				00944-2700-03	2.5	25
				00944-2700-04	5	50
				00944-2700-05	10	100
				00944-2700-06	20	200
				00944-2700-07	30	300
HyQvia 10% (with Recombinant Human Hyaluronidase 160 U/mL)	Baxalta US Inc.	J1575 – Injection, immune globulin/ hyaluronidase, (Hyqvia), 100 mg immune globulin	100 mg	00944-2510-02	2.5	25
				00944-2511-02	5	50
				00944-2512-02	10	100
				00944-2513-02	20	200
				00944-2514-02	30	300
Cuvitru 20%	Baxalta US Inc.	J1555 – Injection, immune globulin (Cuvitru), 100 mg	100 mg	00944-2850-01	1	5
				00944-2850-03	2	10
				00944-2850-05	4	20
				00944-2850-07	8	40
				00944-2850-09	10	50
Cutaquig 16.5%	Octapharma	J1551 – Injection, immune globulin (cutaquig), 100 mg <i>(Effective 07/01/2022)</i>	100 mg	00069-1061-01	1	6
				00069-1802-01	1.65	10
				00069-1476-01	2	12
				00069-1960-01	3.3	20

Drug Name*	Manufacturer	HCP Code	1 Billable unit	NDC	IgG (grams) per SDV	Volume (mL)
		J3590 – unclassified biologics ( <i>Discontinue use on 07/01/2022</i> ) C9399 – unclassified drugs or biologics ( <i>Discontinue use on 07/01/2022</i> )		00069-1509-01	4	24
				00069-1965-01	8	48
Xembify 20%	Grifols	J1558 – Injection, immune globulin (Xembify), 100 mg	100 mg	13533-0810-05	1	5
				13533-0810-10	2	10
				13533-0810-20	4	20
				13533-0810-50	10	50
Immune Globulin, Human, Subcutaneous	N/A	J3590 – unclassified biologics C9399 – unclassified drugs or biologics	N/A	N/A	N/A	N/A

\*90284 – immune globulin (SCIg), human, for use in subcutaneous infusions

## VII. References

- Xembify [package insert]. Research Triangle Park, NC; Grifols Therapeutics, LLC; August 2020. Accessed September 2021.
- Cutaquig [package insert]. Vienna, Austria; Octapharma; October 2021. Accessed October 2021.
- Hizentra [package insert]. Bern, Switzerland; CSL Behring AG; April 2021. Accessed September 2021.
- HyQvia [package insert]. Lexington, MA; Baxalta US Inc.; March 2021. Accessed September 2021.
- Cuvitru [package insert]. Lexington, MA; Baxalta US Inc.; September 2021. Accessed September 2021.
- Gammagard Liquid [package insert]. Lexington, MA; Baxalta US Inc.; March 2021. Accessed September 2021.
- Gamunex®-C [package insert]. Research Triangle Park, NC; Grifols Therapeutics, LLC; January 2020. Accessed September 2021.
- Gammaked [package insert]. Research Triangle Park, NC; Grifols Therapeutics, LLC; January 2020. Accessed September 2021.
- Jeffrey Modell Foundation Medical Advisory Board, 2013. 10 Warning Signs of Primary Immunodeficiency. Jeffrey Modell Foundation, New York, NY
- Orange J, Hossny E, Weiler C, et al. Use of intravenous immunoglobulin in human disease: A review of evidence by members of the Primary Immunodeficiency Committee of the American Academy of Allergy, Asthma and Immunology. *J Allergy Clin Immunol* 2006;117(4 Suppl): S525-53.
- Orange JS, Ballou M, Stiehm, et al. Use and interpretation of diagnostic vaccination in primary immunodeficiency: A working group report of the Basic and Clinical Immunology Interest



Section of the American Academy of Allergy, Asthma & Immunology. *J Allergy Clin Immunol* Vol 130 (3).

12. Bonilla FA, Khan DA, Ballas ZK, et al. Practice Parameter for the diagnosis and management of primary immunodeficiency. *J Allergy Clin Immunol* 2015 Nov;136(5):1186-205.e1-78.
13. Emerson GG, Herndon CN, Sreih AG. Thrombotic complications after intravenous immunoglobulin therapy in two patients. *Pharmacotherapy*. 2002;22:1638-1641.
14. Department of Health (London). Clinical Guidelines for Immunoglobulin Use: Update to Second Edition. August, 2011.
15. Provan, Drew, et al. "Clinical guidelines for immunoglobulin use." Department of Health Publication, London (2008).
16. Dantal J. Intravenous Immunoglobulins: In-Depth Review of Excipients and Acute Kidney Injury Risk. *Am J Nephrol* 2013;38:275-284.
17. Immune Deficiency Foundation. Diagnostic & Clinical Care Guidelines for Primary Immunodeficiency Diseases. 3<sup>rd</sup> Ed. 2015. Avail at: [https://primaryimmune.org/sites/default/files/publications/2015-Diagnostic-and-Clinical-Care-Guidelines-for-PI\\_1.pdf](https://primaryimmune.org/sites/default/files/publications/2015-Diagnostic-and-Clinical-Care-Guidelines-for-PI_1.pdf).
18. Perez EE, Orange JS, Bonilla F, et al. Update on the use of immunoglobulin in human disease: A review of evidence. *J Allergy Clin Immunol*. 2017 Mar;139(3S):S1-S46.
19. Alonso W, Vandenberg P, Lang J, et al. Immune globulin subcutaneous, human 20% solution (Xembify®), a new high concentration immunoglobulin product for subcutaneous administration. *Biologicals*. 2020;64:34-40.
20. Kobayashi RH, Gupta S, Melamed I, et al. Clinical Efficacy, Safety and Tolerability of a New Subcutaneous Immunoglobulin 16.5% (octanorm [cutaquig®]) in the Treatment of Patients with Primary Immunodeficiencies. *Front Immunol*. February 2019 | Volume 10 | Article 40.
21. van Schaik IN, Bril V, van Geloven N, et al. Subcutaneous immunoglobulin for maintenance treatment in chronic inflammatory demyelinating polyneuropathy (CIDP), a multicenter randomised double-blind placebo-controlled trial: the PATH Study. *Lancet Neurol*. 2017;17(1):35-46.
22. Hagan JB, Fasano MB, Spector S, et al. Efficacy and safety of a new 20% immunoglobulin preparation for subcutaneous administration, IgPro20, in patients with primary immunodeficiency. *J Clin Immunol*. 2010;30(5):734-745.
23. Jolles S, Borte M, Nelson R, et al. Long-term efficacy, safety, and tolerability of Hizentra for treatment of primary immunodeficiency disease. *Clin Immunol*. 2014;150(2):161-169.
24. Wasserman RL, Melamed I, Nelson RP Jr, et al. Pharmacokinetics of subcutaneous IgPro20 in patients with primary immunodeficiency. *Clin Pharmacokinet*. 2011;50(6):405-414.
25. Wasserman RL, Melamed I, Kobrynski L, et al. Efficacy, Safety, and Pharmacokinetics of a 10% Liquid Immune Globulin Preparation (GAMMAGARD LIQUID, 10%) Administered Subcutaneously in Subjects with Primary Immunodeficiency Disease. *J Clin Immunol*. 2011 Mar 22. [Epub ahead of print]
26. Food and Drug Administration. Safety, efficacy, and pharmacokinetic studies to support marketing of immune globulin intravenous (human) as replacement therapy for primary humoral immunodeficiency. [https://www.fda.gov/regulatory-information/search-fda-guidance-](https://www.fda.gov/regulatory-information/search-fda-guidance)

documents/safety-efficacy-and-pharmacokinetic-studies-support-marketing-immune-globulin-intravenous-human. Accessed May 28, 2019

27. Wasserman RL, Melamed I, Stein MR, et al; and IGSC, 10% with rHuPH20 Study Group. Recombinant human hyaluronidase-facilitated subcutaneous infusion of human immunoglobulins for primary immunodeficiency. *J Allergy Clin Immunol.* 2012;130(4):951-957.
28. Suez D, Stein M, Gupta S, et al. Efficacy, safety, and pharmacokinetics of a novel human immune globulin subcutaneous, 20% in patients with primary immunodeficiency diseases in North America. *J Clin Immunol.* 2016;36(7):700-712.
29. Roifman CM, Schroeder H, Berger M, et al. Comparison of the efficacy of IGIV-C, 10% (caprylate/chromatography) and IGIV-SD, 10% as replacement therapy in primary immune deficiency: a randomized double-blind trial. *Int Immunopharmacol.* 2003;3(9):1325-1333.
30. Roifman CM, Schroeder H, Berger M, et al, and the IGIV-C in PID Study Group. Comparison of the efficacy of IGIV-C, 10% (caprylate/chromatography) and IGIV-SD, 10% as replacement therapy in primary immune deficiency: a randomized double-blind trial. *Int Immunopharmacol.* 2003;3:1325-1333.
31. Referenced with permission from the NCCN Drugs & Biologics Compendium (NCCN Compendium®) Chronic Lymphocytic Leukemia/Small Lymphocytic Lymphoma, Version 1.2022. National Comprehensive Cancer Network, 2021. The NCCN Compendium® is a derivative work of the NCCN Guidelines®. NATIONAL COMPREHENSIVE CANCER NETWORK®, NCCN®, and NCCN GUIDELINES® are trademarks owned by the National Comprehensive Cancer Network, Inc. To view the most recent and complete version of the Compendium, go online to NCCN.org. Accessed September 2021.
32. Chapel H, Dicato M, Gamm H, et al. Immunoglobulin replacement in patients with chronic lymphocytic leukaemia: a comparison of two dose regimes. *Br J Haematol* 1994 Sep;88(1):209-12. doi: 10.1111/j.1365-2141.1994.tb05002.x.

## Appendix 1 – Covered Diagnosis Codes (All Products)

ICD-10	ICD-10 Description
C91.10	Chronic lymphocytic leukemia of B-cell type not having achieved remission
C91.11	Chronic lymphocytic leukemia of B-cell type in remission
C91.12	Chronic lymphocytic leukemia of B-cell type in relapse
D80.0	Hereditary hypogammaglobulinemia
D80.1	Nonfamilial hypogammaglobulinemia
D80.2	Selective deficiency of immunoglobulin A [IgA]
D80.3	Selective deficiency of immunoglobulin G [IgG] subclasses
D80.4	Selective deficiency of immunoglobulin M [IgM]
D80.5	Immunodeficiency with increased immunoglobulin M [IgM]
D80.7	Transient hypogammaglobulinemia of infancy
D81.0	Severe combined immunodeficiency [SCID] with reticular dysgenesis
D81.1	Severe combined immunodeficiency [SCID] with low T- and B-cell numbers

ICD-10	ICD-10 Description
D81.2	Severe combined immunodeficiency [SCID] with low or normal B-cell numbers
D81.6	Major histocompatibility complex class I deficiency
D81.7	Major histocompatibility complex class II deficiency
D81.89	Other combined immunodeficiencies
D81.9	Combined immunodeficiency, unspecified
D82.0	Wiskott-Aldrich syndrome
D83.0	Common variable immunodeficiency with predominant abnormalities of B-cell numbers and function
D83.2	Common variable immunodeficiency with autoantibodies to B- or T-cells
D83.8	Other common variable immunodeficiencies
D83.9	Common variable immunodeficiency, unspecified

### Additional covered diagnosis codes applicable to Hizentra ONLY:

ICD-10	ICD-10 Description
G61.81	Chronic inflammatory demyelinating polyneuritis
G61.89	Other inflammatory polyneuropathies
G62.89	Other specified polyneuropathies

### 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), Local Coverage Determinations (LCDs), and Local Coverage Articles (LCAs) may exist and compliance with these policies is required where applicable. They can be found at: <https://www.cms.gov/medicare-coverage-database/search.aspx>. Additional indications may be covered at the discretion of the health plan.

Medicare Part B Covered Diagnosis Codes (applicable to existing NCD/LCD/LCA): N/A

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 Corp (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 Corp (WPS)
N (9)	FL, PR, VI	First Coast Service Options, Inc.
J (10)	TN, GA, AL	Palmetto GBA, LLC

**Medicare Part B Administrative Contractor (MAC) Jurisdictions**

<b>Jurisdiction</b>	<b>Applicable State/US Territory</b>	<b>Contractor</b>
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