HIGHMARK HEALTH OPTIONS

Updated: 04/2025
DMMA Approved: 05/2025

Request for Prior Authorization for Gene Therapy Agents
Website Form – www.highmarkhealthoptions.com
Submit request via: Fax - 1-855-476-4158

All requests for Gene Therapy Agents without their own policy require a prior authorization and will be screened for medical necessity and appropriateness using the criteria listed below.

Gene Therapy Agents Prior Authorization Criteria:
Gene therapies include Kebilidi (eladocagene exuparvovec, Zynteglo (betibeglogene autotemcel), Skysona (elivaldogene autotemcel), Hemgenix (etranacogene dezaparvovec), Roctavian (valoctocogene roxaparvovec), Elevidys (delandistrogene moxeparvovec-rokl), Lenmeldy (atidarsagene autotemcel), Casgevy (exagamglogene autotemcel), Lyfgenia (Lovotibeglogene autotemcel) and Adstiladrin (nadofaragene firadenovec-vncg). New products with this classification will require the same documentation.

For all requests the following criteria must be met in addition to the diagnosis specific criteria below:

- Is prescribed for an indication that is included in the U.S. Food and Drug Administration (FDA)-approved package labeling OR a medically accepted indication
- The requested dose and frequency is in accordance with FDA-approved labeling, nationally recognized compendia, and/or evidence-based practice guidelines
- Must be age-appropriate according to FDA-approved labeling, nationally recognized compendia, or evidence-based practice guidelines
- The member is not currently enrolled in a clinical trial for the requested drug or has previously received the requested gene therapy or any other gene therapy.
- Does not have any contraindications to the requested medication

For Hemgenix (etranacogene dezaparvovec) requests:

Coverage may be provided with a diagnosis of Hemophilia B (congenital Factor IX deficiency) and the following criteria is met:

- Member must have severe or moderately severe hemophilia B (congenital factor IX deficiency) defined as equal to or less than 2% of normal circulating factor IX confirmed by blood coagulation testing
- Must have baseline liver function tests assessed prior to and after therapy for at least three months and be within normal range
- Members with preexisting risk factors for hepatocellular carcinoma (e.g., members with cirrhosis, advanced hepatic fibrosis, hepatitis C or B, non-alcoholic fatty liver disease (NAFLD), chronic alcohol consumption, non-alcoholic steatohepatitis (NASH), and advanced age) must have abdominal ultrasound screenings and be monitored regularly (e.g., annually) for alpha-fetoprotein (AFP) elevations following administration
- Is prescribed by a hematologist or hemophilia treatment center practitioner
- Member has received IX prophylactic or on-demand replacement therapy for ≥ 150 accumulated days and is currently using factor IX prophylaxis therapy

- Member has ≥ 12 bleeding episodes if receiving on-demand therapy over the preceding 12 months. Does not apply to patients on prophylaxis.
- Member must have a baseline anti-AAV5 antibody titer of ≤ 1:678 measured by ELISA
- Member must not have any of the following:
  - Inhibitor antibodies to factor IX
  - A positive HIV test during time of screening that is not controlled with anti-viral therapy
  - Active infection with hepatitis B or C virus at screening
  - History of hepatitis B or C exposures, currently controlled by antiviral therapy
  - Prior hemophilia AAV-vector based gene therapy
- **Duration of Approval:** One lifetime dose

For Roctavian (valoctocogene roxaparvovec) requests:

Coverage may be provided with a diagnosis of Hemophilia A (congenital Factor VIII deficiency) and the following criteria is met:

- Member must have severe hemophilia A (congenital factor VIII deficiency) defined as less than 1% of normal circulating factor VIII confirmed by blood coagulation testing
- Member must not have any pre-existing antibodies to adeno-associated virus serotype 5 detected by an FDA approved test.
- Member must not have any contraindications to receiving therapy such as active infections (either acute or uncontrolled chronic), significant hepatic fibrosis (stage 3 or 4) or cirrhosis or a known hypersensitivity to mannitol.
- Member meets both of the following:
  - No previous documented history of a detectable FVIII inhibitor
  - Member has inhibitor level assay < 1 Bethesda units (BU) on 2 consecutive occasions at least one week apart within the last 12 months
- Must have baseline liver function tests assessed prior to and after therapy for at least three months and be within normal range
- Members with preexisting risk factors for hepatocellular carcinoma (e.g., members with cirrhosis, advanced hepatic fibrosis, hepatitis C or B, non-alcoholic fatty liver disease (NAFLD), chronic alcohol consumption, non-alcoholic steatohepatitis (NASH), and advanced age) must have abdominal ultrasound screenings and be monitored regularly (e.g., annually) for alpha-fetoprotein (AFP) elevations following administration
- Is prescribed by a hematologist or hemophilia treatment center practitioner
- Member has received VIII prophylactic or on-demand replacement therapy for ≥ 150 accumulated days and is currently using factor VIII prophylaxis therapy
- Member has ≥ 12 bleeding episodes if receiving on-demand therapy over the preceding 12 months. Does **not** apply to patients on prophylaxis.
- Member must not have any of the following:
  - A positive HIV test during time of screening that is not controlled with anti-viral therapy
  - Active infection with hepatitis B or C virus at screening
  - History of hepatitis B or C exposures, currently controlled by antiviral therapy

  - Prior hemophilia AAV-vector based gene therapy
- **Duration of Approval:** One lifetime dose

For Elevidys (delandistrogene moxeparvovec-rokl) requests:

Coverage may be provided with a diagnosis of Duchenne muscular dystrophy (DMD) and the following criteria is met:

- A confirmed diagnosis of DMD by submission of lab testing demonstrating mutation of the dystrophin (DMD) gene by either:
  - A confirmed frameshift mutation OR
  - A premature stop codon mutation between exons 18 to 58 in the DMD gene
- Member meets either of the following criteria:
  - Member is ambulatory (e.g., able to walk with or without assistance, not wheelchair dependent).
  - Member is non-ambulatory and has a Performance Upper Limb (PUL) entry item score of at least 3 and a total PUL score of 20 – 40.
- The member must not have any deletion in exon 8 and/or exon 9 in the DMD gene
- Member does not have signs of cardiomyopathy (e.g., ejection fraction < 40%)
- Member does not currently have an active infection
- The member must be on a stable dose of corticosteroids for DMD for at least 12 weeks prior to therapy unless contraindicated
- The member must have a baseline anti-AAVrh74 antibody titers <1:400 measured by ELISA
- Must be prescribed by or in consultation with a neurologist who has experience in the treatment and management of DMD
- Elevidys will not be used in combination with exon-skipping therapies (e.g., casimersen, eteplirsen, golodirsen, viltolarsen).
- There is documentation of a baseline evaluation, including a standardized assessment of motor function, by a neurologist with experience treating DMD and liver function, platelet count, and troponin-I levels have been assessed at baseline and will be monitored as clinically appropriate.
- **Duration of Approval**: One lifetime dose

For Zynteglo (betibeglogene autotemcel) requests:

Coverage may be provided with a diagnosis of beta-thalassemia and the following criteria is met:

- The member must be transfusion-dependent β-thalassaemia (TDT) who do not have a β0 /β0 genotype, for whom haematopoietic stem cell (HSC) transplantation is appropriate but a human leukocyte antigen (HLA)-matched related HSC donor is not available
- Members are considered to be transfusion-dependent if they had a history of transfusions of at least 100 mL/kg/year of RBCs or with ≥8 transfusions of RBCs per year in the 2 years preceding enrolment.
- Is prescribed by a hematologist, stem cell transplantation specialist or in the treatment of members with TDT
- Must be administered in a qualified treatment center

- Physician must confirm that HSC transplantation is appropriate for the member before myeloablative conditioning is initiated
- Member must not have had previous treatment with HSC gene therapy
- Member must not be pregnant or breast-feeding
- All patients should be tested for HIV prior to mobilization and apheresis to ensure acceptance of the apheresis material for manufacturing
- **Duration of Approval:** One lifetime dose

For Skysona (elivaldogene autotemcel) requests:

Coverage may be provided with a diagnosis of **cerebral adrenoleukodystrophy (CALD)** and the following criteria is met:

- Must have early, active CALD defined by:
  - Elevated very long chain fatty acids (VLCFA) values
  - Active CNS disease established by central radiographic review of brain magnetic resonance imaging (MRI)
  - Loes score between 0.5 and 9
  - Gadolinium enhancement (GdE+) on MRI of demyelinating lesions
  - Neurologic function score (NFS) of ≤ 1 demonstrating asymptomatic or mild disease
- Member must have confirmed mutations in the ABCD1 gene
- Must be prescribed by a neurologist or ALD specialist.
- The requested dose and frequency is in accordance with FDA-approved labeling, nationally recognized compendia, and/or evidence-based practice guidelines
- Skysona should not be administered in members with active infections.
- Member must have a negative serology test for HIV.
- Member must not have been a recipient of an allogenic transplant or gene therapy
- **Duration of Approval:** One lifetime dose

For Casgevy (exagamglogene autotemcel) requests:

Coverage may be provided with a diagnosis of **severe sickle cell disease (SCD)** and the following criteria is met:

- Diagnosis is confirmed by electrophoresis demonstrating the presence of sickle cell disease (HbSS, HbSC, HbSβ^0-thalassemia, or HbSβ^+-thalassemia).
- Documentation the member is a candidate for an allogenic hematopoietic stem cell transplantation (HSCT) but ineligible due to absence of an appropriate donor (i.e., individual does not have a human leukocyte antigen (HLA)-matched donor; or individual has an HLA-matched donor, but the potential donor is not able or is not willing to donate)
- Documentation the member is currently on a disease modifying therapies for SCD (e.g., hydroxyurea, crizanlizumab, voxelotor) and will discontinue use 8 weeks before the planned start of mobilization and conditioning

- While receiving appropriate standard treatment for SCD, individual had at least four severe vaso-occlusive crises or events (VOCs/VOEs) in the previous 2 years, which can be defined by one or more of the following (i, ii, iii, iv, or v):
    i. An episode of acute pain that resulted in a visit to a medical facility that required administration of at least ONE of the following (a or b):
        a) Intravenous opioid; OR
        b) Intravenous nonsteroidal anti-inflammatory drug;
    ii. Acute chest syndrome (acute chest syndrome is defined by the presence of a new pulmonary infiltrate associated with pneumonia-like symptoms (e.g., chest pain, fever [> 99.5°F], tachypnea, wheezing or cough, or findings upon lung auscultation);
    iii. Acute hepatic sequestration (defined by a sudden increase in liver size associated with pain in the right upper quadrant, abnormal results of liver function test not due to biliary tract disease, and the reduction of hemoglobin concentration by ≥2 g/dL below the baseline value)
    iv. Acute splenic sequestration (defined by an enlarged spleen, left upper quadrant pain, and an acute decrease in hemoglobin concentration of ≥ 2 g/dL below the baseline value);
    v. Acute priapism lasting >2 hours and requiring a visit to a medical facility
- Must be prescribed by a hematologist/oncologist or stem cell transplant physician
- Member must not have:
    - Advanced liver disease
    - History of untreated Moyamoya disease or presence of Moyamoya disease
- For members who are 12-18 years of age, members must have normal transcranial Doppler (TCD)
- Member does not currently have an active bacterial, viral, fungal or parasitic infection.
- Member must not be pregnant or breast-feeding
- Member must not need curative anticoagulation therapy during the period of conditioning through platelet engraftment
- Member must not have been a recipient of an allogenic hematopoietic cell transplant or previous gene therapy
- Member must not have any of the following:
    - Advanced liver disease (e.g., alanine transaminase > 3 times upper limit of normal; direct bilirubin value > 2.5 times upper limit of normal; baseline prothrombin time (international normalized ratio [INR]) > 1.5 times upper limit of normal; cirrhosis; bridging fibrosis; or active hepatitis);
    - Severe cerebral vasculopathy as defined by history of untreated Moyamoya disease or presence of Moyamoya disease that puts the individual at risk of bleeding, per the prescribing physician;
    - Prior or current malignancy, myeloproliferative disorder, or significant immunodeficiency disorder;
- All members must be tested for HIV-1, HIV-2, HBV, HCV and Human T-lymphotropic virus-1 and -2 prior to mobilization and apheresis to ensure acceptance of the apheresis material for manufacturing
- **Duration of Approval:** One lifetime dose

For Casgevy (exagamglogene autotemcel) requests:

Coverage may be provided with a diagnosis of beta-thalassemia and the following criteria is met:

- Documentation the member has transfusion-dependent β-thalassemia (TDT) confirmed by genetic testing with one of the following genotypes:
  - Non-β0/β0 genotype (e.g., β0/β+, βE/β0, and β+/β+); OR
  - β0/β0 genotypes (other examples include β0/β+(IVS-I-110) and β+(IVS-I-110)/β+(IVS-I-110);
- The member must be transfusion depended defined as:
  - History of transfusions of ≥100 mL per kg of body weight of packed red blood cells per year in the previous 2 years; OR
  - History of transfusions of ≥10 units of packed red blood cells per year in the previous 2 years
- Documentation that the member is a candidate for an allogeneic HSCT, but ineligible due to absence of an appropriate donor (i.e., individual does not have a Human Leukocyte Antigen (HLA)-matched donor; or individual has an HLA-matched donor, but the potential donor is not able or is not willing to donate);
- Must be prescribed by a hematologist/oncologist or stem cell transplant physician
- Must be administered in a qualified treatment center
- Member must not have any of the following:
  - Severely elevated iron in the heart (i.e. cardiac T2* less than 10 msec by magnetic resonance imaging [MRI] or left ventricular ejection fraction [LVEF] < 45% by echocardiogram)
  - Advanced liver disease (aspartate transaminase [AST] or alanine transaminase [ALT] > 3 × the upper limit of normal [ULN], or direct bilirubin value > 2.5 × ULN, or if a liver biopsy demonstrated bridging fibrosis or cirrhosis [liver biopsy was performed if liver iron content was ≥ 15 mg/g by MRI])
  - An available 10/10 human leukocyte antigen matched related hematopoietic stem cell donor
  - Associated α-thalassemia and >1 alpha deletion or alpha multiplications
  - Sickle cell beta thalassemia variant
  - Clinically significant and active bacterial, viral, fungal, or parasitic infection
  - White blood cell (WBC) count <3 × 10^9/L or platelet count <50 × 10^9/L not related to hypersplenism
  - Severe cerebral vasculopathy as defined by history of untreated Moyamoya disease or presence of Moyamoya disease that puts the individual at risk of bleeding, per the prescribing physician
  - Prior or current malignancy, myeloproliferative disorder, or significant immunodeficiency disorder
- Member must not have had previous treatment with a hematopoietic stem cell (HSC) gene therapy or prior allo-HSCT
- Member must be able to receive a red blood cell transfusion
- Documentation the member will have discontinued iron chelation therapy for at least 7 days prior to myeloablative conditioning (e.g., iron chelators used for this condition include deferoxamine injection, deferiprone tablets or solution, and deferasirox tablets)
- Member does not currently have an active bacterial, viral, fungal or parasitic infection.

- Member must not be pregnant or breast-feeding
- All members must be tested for HIV-1, HIV-2, HBV, HCV and Human T-lymphotropic virus-1 and -2 prior to mobilization and apheresis to ensure acceptance of the apheresis material for manufacturing
- **Duration of Approval:** One lifetime dose

For Lyfgenia (lovotibeglogene autotemcel) requests:

Coverage may be provided with a diagnosis of **severe sickle cell disease (SCD)** and the following criteria is met:

- Diagnosis is confirmed by electrophoresis demonstrating the presence of sickle cell disease with either βS/βS or βS/β0 or βS/β+ genotype
- Documentation the member is a candidate for an allogenic hematopoietic stem cell transplantation (HSCT) but ineligible due to absence of an appropriate donor (i.e., individual does not have a human leukocyte antigen (HLA)-matched donor; or individual has an HLA-matched donor, but the potential donor is not able or is not willing to donate)
- Documentation the member is currently on a disease modifying therapies for SCD (e.g., hydroxyurea, crizanlizumab, voxelotor) and will discontinue use 8 weeks before the planned start of mobilization and conditioning
- While receiving appropriate standard treatment for SCD, individual had at least four severe vaso-occlusive crises or events (VOCs/VOEs) in the previous 2 years, which can be defined by one or more of the following (i, ii, iii, iv, or v):
    i. An episode of acute pain that resulted in a visit to a medical facility that required administration of at least ONE of the following (a or b):
        a) Intravenous opioid; OR
        b) Intravenous nonsteroidal anti-inflammatory drug;
    ii. Acute chest syndrome (acute chest syndrome is defined by the presence of a new pulmonary infiltrate associated with pneumonia-like symptoms (e.g., chest pain, fever [> 99.5°F], tachypnea, wheezing or cough, or findings upon lung auscultation);
    iii. Acute hepatic sequestration (defined by a sudden increase in liver size associated with pain in the right upper quadrant, abnormal results of liver function test not due to biliary tract disease, and the reduction of hemoglobin concentration by ≥2 g/dL below the baseline value)
    iv. Acute splenic sequestration (defined by an enlarged spleen, left upper quadrant pain, and an acute decrease in hemoglobin concentration of ≥ 2 g/dL below the baseline value);
    v. Acute priapism lasting >2 hours and requiring a visit to a medical facility
- Must be prescribed by a hematologist/oncologist or stem cell transplant physician
- Member must have a Karnofsky performance status of ≥ 60 (≥16 years of age) or a Lansky performance status of ≥60 (<16 years of age).
- The member has experienced, at any time in the past, an inadequate response, intolerance or contraindication to a trial of hydroxyurea
- Member must not have any of the following:
    - Advanced liver disease (e.g., alanine transaminase > 3 times upper limit of normal; direct bilirubin value > 2.5 times upper limit of normal; baseline prothrombin time

(international normalized ratio [INR]) > 1.5 times upper limit of normal; cirrhosis; bridging fibrosis; or active hepatitis);
    - Severe cerebral vasculopathy as defined by history of untreated Moyamoya disease or presence of Moyamoya disease that puts the individual at risk of bleeding, per the prescribing physician;
    - Prior or current malignancy, myeloproliferative disorder, or significant immunodeficiency disorder;
- For members who are 12-18 years of age, members must have normal transcranial Doppler (TCD)
- Member must not need curative anticoagulation therapy during the period of conditioning through platelet engraftment
- Member must not be pregnant or breast-feeding
- Member must be able to receive a red blood cell transfusion
- All members must be tested for HIV-1, HIV-2, HBV, HCV and Human T-lymphotropic virus-1 and -2 prior to mobilization and apheresis to ensure acceptance of the apheresis material for manufacturing
- Member does not currently have an active bacterial, viral, fungal or parasitic infection.
- Member must not have been a recipient of an allogenic hematopoietic cell transplant or previous gene therapy
- **Duration of Approval:** One treatment per lifetime

For Lenmeldy (atidarsagene autotemcel) requests:

Coverage may be provided with a diagnosis of **metachromatic leukodystrophy (MLD)** and the following criteria is met:

- Member must have one of the following:
    - Pre-symptomatic late infantile (PSLI) MLD
    - Pre-symptomatic early juvenile (PSEJ) MLD
    - Early symptomatic early juvenile (ESEJ) MLD
- A confirmed diagnosis by **all** of the following:
    - Biochemical testing documenting human arylsulfatase A (ARSA) gene activity is below the normal range for the laboratory performing the test
    - The presence of two disease-causing ARSA alleles, either known or novel mutations, identified on genetic testing
    - If novel mutations are identified, a 24-hour urine collection showing elevated sulfatide levels
- Must be prescribed by or in consultation with a physician who specializes in the treatment of MLD
- The member does not have evidence of residual cells of donor origin if the member has received a prior allogeneic hematopoietic stem cell transplant (allo-HSCT).
- **Duration of Approval:** One lifetime dose

For Kebilidi (eladocagene exuparvovec-tneq) requests:

Coverage may be provided with a diagnosis of **aromatic L amino acid decarboxylase (AADC) deficiency** and the following criteria is met:

- Diagnosis must be confirmed based on **all** of the following:
  - Genetic testing showing biallelic mutations in the DOPA decarboxylase (DDC) gene
  - Reduced levels of 5-hydroxyindoleacetic acid (5-HIAA), homovanillic acid (HVA) and 3-methoxy-4hydroxyphenylglycol (MHPG) and high concentrations of 3-O-methyldopa (3-OMD), L-Dopa, and 5OH tryptophan (5-HTP) in the cerebral spinal fluid (CSF)
  - Reduced aromatic L-amino acid decarboxylase (AADC) activity in the plasma
- Must be prescribed by or in consultation with a pediatric neurologist
- Must present with classical clinical characteristics of AADC deficiency, such as oculogyric crises, hypotonia, and developmental delay
- Must not have any significant structural brain abnormality
- Must not have an anti-AAV2 neutralizing antibody titer over 1,200 folds
- Member must not have received prior treatment with any other AAV2-based gene therapy despite indication or are being considered for treatment with any other AAV2-based gene therapy
- **Duration of Approval**: One lifetime dose

For Adstiladrin (nadofaragene firadenovec-vncg) requests:
Coverage may be provided with a diagnosis of **non-muscle invasive bladder cancer (NMIBC) with carcinoma in situ (CIS)** and the following criteria is met:

- The member has a confirmed diagnosis of NMIBC with CIS with or without papillary tumors
- The member's disease is high-risk and BCG unresponsive defined as:
  - Persistent disease following adequate BCG therapy,
  - Disease recurrence after an initial tumor-free state following adequate BCG therapy, or
  - T1 disease following a single induction course of BCG
- The member is not immunocompromised or immunodeficient
- Member must not have extra-vesical (i.e., urethra, ureter, or renal pelvis), muscle invasive (T2-T4), or metastatic urothelial carcinoma
- The member is ineligible for or has elected not to undergo cystectomy
- Member must have an ECOG performance status ≤2
- **Initial Duration of Approval:** 6 months
- **Reauthorization criteria**
  - The member continues to meet the requirements for initial therapy and has been treated with Adstiladrin (nadofaragene firadenovec-vncg) with no adverse reactions.
  - The member has no signs of unacceptable toxicity (such as risk of disseminated adenovirus infection) while on treatment with Adstiladrin (nadofaragene firadenovec-vncg).
- **Reauthorization Duration of Approval: 6 months**

Coverage may be provided for any non-FDA labeled indication if it is determined that the use is a medically accepted indication supported by nationally recognized pharmacy compendia or peer-

reviewed medical literature for treatment of the diagnosis(es) for which it is prescribed. These requests will be reviewed on a case by case basis to determine medical necessity.

Drugs are authorized in generic form unless the branded product is on the preferred drug list or the prescriber has indicated in writing that the branded product is medically necessary. If only the branded product is on the preferred drug list, the generic form will be considered non-preferred and shall not require the prescriber to indicate in writing that the branded product is medically necessary.

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GENE THERAPY AGENTS
PRIOR AUTHORIZATION FORM- PAGE 1 of 4

Please complete and fax all requested information below including any progress notes, laboratory test results, or chart documentation as applicable to Highmark Health Options Pharmacy Services. **FAX: (855) 476-4158**
If needed, you may call to speak to a Pharmacy Services Representative. **PHONE**: (844) 325-6251 Mon – Fri 8:00 am to 7:00 pm

PROVIDER INFORMATION

Requesting Provider:	NPI:
Provider Specialty:	Office Contact:
Office Address:	Office Phone:
	Office Fax:

MEMBER INFORMATION

Member Name:	DOB:	
Member ID:	Member weight:	Height:

REQUESTED DRUG INFORMATION

Medication:	Strength:	
Directions:	Quantity:	Refills:
Is the member currently receiving requested medication? ☐ Yes ☐ No	Date Medication Initiated:	
Is this medication being used for a chronic or long-term condition for which the medication may be necessary for the life of the patient? ☐ Yes ☐ No		

Billing Information

This medication will be billed: ☐ at a pharmacy **OR** ☐ medically, JCODE:_____
Place of Service: ☐ Hospital ☐ Provider's office ☐ Member's home ☐ Other

Place of Service Information

Name:	NPI:
Address:	Phone:

MEDICAL HISTORY (Complete for ALL requests)

Diagnosis:	ICD Code:

Hemophilia A:

Does the member have severe hemophilia A? ☐ Yes, normal factor activity level:_____ ☐ No
Does the member have any pre-existing antibodies to adeno-associated virus serotype 5 detected by an FDA approved test? ☐ Yes ☐ No
Does the member have any contraindications to receiving therapy? ☐ Yes ☐ No
Did the member have baseline liver function tests assessed prior to therapy and was it within normal range? ☐ Yes ☐ No
Will the member have liver function testing done for at least 3 months after therapy? ☐ Yes ☐ No
Did the member have abdominal ultrasound screenings if they have preexisting risk factors for hepatocellular carcinoma? ☐ Yes ☐ No
Has the member had any documented history of a detectable FVIII inhibitor or an inhibitor level assay <1 BU on 2 consecutive occasions at least one week apart with the last 12 months? ☐ Yes, please explain below. ☐ No
Has the member had ≥ 12 bleeding episodes if receiving on-demand therapy over the preceding 12 months? Does **not** apply to patients on prophylaxis. ☐ Yes ☐ No
Has the member received FVIII prophylactic or on-demand replacement therapy for ≥ 150 accumulated days and still on current therapy? ☐ Yes ☐ No
Does the member have a positive HIV test or active infection with Hepatitis B or C? ☐ Yes ☐ No
Has the member had prior hemophilia AAV-vector based gene therapy? ☐ Yes ☐ No

Hemophilia B:

Does the member have severe or moderately severe B? ☐ Yes, normal factor activity level:_____ ☐ No
Did the member have baseline liver function tests assessed prior to therapy and was it within normal range? ☐ Yes ☐ No
Will the member have liver function testing done for at least 3 months after therapy? ☐ Yes ☐ No
Did the member have abdominal ultrasound screenings if they have preexisting risk factors for hepatocellular carcinoma? ☐ Yes ☐ No

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GENE THERAPY AGENTS
PRIOR AUTHORIZATION FORM (CONTINUED) – PAGE 2 OF 4

Please complete and fax all requested information below including any progress notes, laboratory test results, or chart documentation as applicable to Highmark Health Options Pharmacy Services. **FAX: (855) 476-4158**
If needed, you may call to speak to a Pharmacy Services Representative. **PHONE**: (844) 325-6251 Mon – Fri 8 am to 7 pm

MEMBER INFORMATION

Member Name:	DOB:	
Member ID:	Member weight:	Height:

MEDICAL HISTORY (Complete for ALL requests)

Diagnosis:	ICD Code:

Hemophilia B (continued):

Has the member had ≥ 12 bleeding episodes if receiving on-demand therapy over the preceding 12 months? Does **not** apply to patients on prophylaxis. ☐ Yes ☐ No

Has the member received IX prophylactic or on-demand replacement therapy for ≥ 150 accumulated days and still on current therapy? ☐ Yes ☐ No

What is the members baseline anti-AAV5 antibody titer measured by ELISA? _____

Does the member have inhibitor antibodies to factor IX? ☐ Yes ☐ No

Does the member have a positive HIV test or active infection with Hepatitis B or C? ☐ Yes ☐ No

Has the member had prior hemophilia AAV-vector based gene therapy? ☐ Yes ☐ No

DMD:

Does the member have a diagnosis of DMD confirmed by submission of lab testing demonstrating mutation of the dystrophin (DMD) gene by either a confirmed frameshift mutation OR a premature stop codon mutation between exons 18 to 58 in the DMD gene? ☐ Yes ☐ No

Is the member ambulatory? ☐ Yes ☐ No

Is the member non-ambulatory? ☐ Yes ☐ No If yes, what is the member's total PUL score? _____

Does the member have any deletion in exon 8 and/or exon 9 in the DMD gene? ☐ Yes ☐ No

Does the member have cardiomyopathy with an ejection fraction less than 40%? ☐ Yes ☐ No

Does the member currently have an active infection? ☐ Yes ☐ No

Is the member on a stable dose of corticosteroids for DMD for at least 12 weeks prior to therapy? ☐ Yes ☐ No

What is the member's baseline anti-AAVrh74 antibody titers level measured by ELISA? _____

Is the requested medication being used in combination with exon-skipping therapies? ☐ Yes ☐ No

Is there documentation of a baseline evaluation including a standardized assessment of motor function done by a neurologist with experience in treating DMD and liver function, platelet count and troponin-I levels assessed to be clinically appropriate? ☐ Yes ☐ No

Beta-Thalassemia:

Is the member transfusion-dependent β-thalassaemia (TDT) who does not have a β0 /β0 genotype, for whom haematopoietic stem cell (HSC) transplantation is appropriate but a human leukocyte antigen (HLA)-matched related HSC donor is not available? ☐ Yes ☐ No

Is the member considered transfusion-dependent? ☐ Yes ☐ No

Is the medication being administered in a qualified treatment center? ☐ Yes ☐ No

Has the physician confirmed that HSC transplantation is appropriate for the member before myeloablative conditioning is initiated? ☐ Yes ☐ No

Does the member have any contraindications to requested therapy? ☐ Yes ☐ No

Has the member had previous treatment with HSC gene therapy? ☐ Yes ☐ No

Is the member pregnant or breast-feeding? ☐ Yes ☐ No

Has the member been tested for HIV prior to mobilization and apheresis to ensure acceptance of the apheresis material for manufacturing? ☐ Yes ☐ No

CALD:

Does the member have early, active CALD? ☐ Yes ☐ No

Does the member have elevated VLCFA? ☐ Yes ☐ No Value: _____

Has the member had an MRI establishing active CNS disease with GdE+ of demyelinating lesions? ☐ Yes ☐ No

What is the Loes score? _____

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GENE THERAPY AGENTS
PRIOR AUTHORIZATION FORM (CONTINUED) – PAGE 3 OF 4

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MEMBER INFORMATION

Member Name:	DOB:	
Member ID:	Member weight:	Height:

MEDICAL HISTORY (Complete for ALL requests)

Diagnosis:	ICD Code:

CALD (continued):
What is the NFS score? _____
Does the member have confirmed mutations in the ABCD1 gene? ☐ Yes ☐ No
Does the member have an active infection? ☐ Yes ☐ No
Does the member have HIV? ☐ Yes ☐ No
Has the member received an allogenic transplant or gene therapy previously? ☐ Yes ☐ No

Sickle Cell Disease (SCD):
For Casgevy only:
Does the member have severe sickle cell disease confirmed by electrophoresis? ☐ Yes, what is the genotype? _____ ☐ No
Is the member eligible for a hematopoietic stem cell transplant but a stem cell donor is not available? ☐ Yes ☐ No
Does the member have a history of at least 2 severe vaso-occlusive crisis (VOC) events during the past 24 months? ☐ Yes, please list the dates:_____ ☐ No
Does the member have advanced liver disease or a history of untreated Moyamoya disese or the presence of Moyamoya disease? ☐ Yes ☐ No
For members who are 12-18 years of age, does the member have a normal transcranial Doppler (TCD)? ☐ Yes ☐ No
If the member is currently on disease modifying therapies for SCD (e.g., hydroxyurea, crizanlizumab, voxelotor), have they discontinued the product at least 8 weeks prior to start of mobilization and conditioning? ☐ Yes ☐ No
Does the member have clinically significant and active bacterial, viral, fungal or parasitic infection? ☐ Yes ☐ No
Has the member been a recipient of an allogenic transplant or gene therapy previously? ☐ Yes ☐ No

For Lyfgenia only:
Does the member have severe sickle cell disease confirmed by electrophoresis? ☐ Yes, what is the genotype? _____ ☐ No
Is the member eligible for a hematopoietic stem cell transplant but a stem cell donor is not available? ☐ Yes ☐ No
Does the member have a history of at least 4 severe vaso-occlusive events (VOE) during the past 24 months? ☐ Yes, please list the dates:_____ ☐ No
Does the member have a Karnofsky performance status of ≥ 60 (≥16 years of age) or a Lansky performance status of ≥60 (<16 years of age)? ☐ Yes ☐ No
Has the member tried and failed hydroxyurea (HU) at any point in the past or had an intolerance to HU? ☐ Yes ☐ No
Does the member have advanced liver disease or a history of untreated Moyamoya disese or the presence of Moyamoya disease? ☐ Yes ☐ No
For members who are 12-18 years of age, does the member have a normal transcranial Doppler (TCD)? ☐ Yes ☐ No
Does the member need curative anticoagulation therapy during the period of conditioning through platelet engraftment?
Is the member able to receive a red blood cell transfusion? ☐ Yes ☐ No
Does the member have a negative serology test for HIV? ☐ Yes ☐ No
Does the member have clinically significant and active bacterial, viral, fungal or parasitic infection? ☐ Yes ☐ No
Has the member been a recipient of an allogenic transplant or gene therapy previously? ☐ Yes ☐ No

Metachromatic Leukodystrophy (MLD):
Does the member have one of the following (please select the appropriate one): ☐ Yes ☐ No ☐ Other:_____
☐ Pre-symptomatic late infantile (PSLI) MLD
☐ Pre-symptomatic early juvenile (PSEJ) MLD
☐ Early symptomatic early juvenile (ESEJ) MLD

HIGHMARK HEALTH OPTIONS

Updated: 04/2025
DMMA Approved: 05/2025

GENE THERAPY AGENTS
PRIOR AUTHORIZATION FORM (CONTINUED) – PAGE 4 OF 4

Please complete and fax all requested information below including any progress notes, laboratory test results, or chart documentation as applicable to Highmark Health Options Pharmacy Services. **FAX: (855) 476-4158**
If needed, you may call to speak to a Pharmacy Services Representative. **PHONE**: (844) 325-6251 Mon – Fri 8 am to 7 pm

MEMBER INFORMATION

Member Name:	DOB:	
Member ID:	Member weight:	Height:

MEDICAL HISTORY (Complete for ALL requests)

Diagnosis:	ICD Code:

MLD continued:

Has the diagnosis been confirmed by all of the following (submit confirmatory laboratory results): biochemical testing documenting human arylsulfatase A (ARSA) gene activity is below the normal range for the laboratory performing the test, the presence of two disease-causing ARSA alleles, either known or novel mutations, identified on genetic testing, and if novel mutations are identified, a 24-hour urine collection showing elevated sulfatide levels? ☐ Yes ☐ No

Is the requested medication being prescribed by or in consultation with a physician who specializes in the treatment of MLD? ☐ Yes ☐ No

Is there evidence of residual cells of donor origin if the member has received a prior allogeneic hematopoietic stem cell transplant (allo-HSCT)? ☐ Yes ☐ No

Aromatic L amino acid decarboxylase (AADC) deficiency:

Has the diagnosis been confirmed based on **all** of the following (submit confirmatory laboratory results): ☐ Yes ☐ No

- Genetic testing, reduced levels of 5-hydroxyindoleacetic acid (5-HIAA), homovanillic acid (HVA) and 3-methoxy-4hydroxyphenylglycol (MHPG) and high concentrations of 3-O-methyldopa (3-OMD), L-Dopa, and 5OH tryptophan (5-HTP) in the cerebral spinal fluid (CSF) AND reduced aromatic L-amino acid decarboxylase (AADC) activity in the plasma

Is the requested medication being prescribed by or in consultation with a pediatric neurologist? ☐ Yes ☐ No

Does the member present with classical clinical characteristics of AADC deficiency, such as oculogyric crises, hypotonia, and developmental delay? ☐ Yes ☐ No

Does the member have any significant structural brain abnormality? ☐ Yes ☐ No

Does the member have anti-AAV2 neutralizing antibody titer over 1,200 folds? ☐ Yes ☐ No

Has the member received prior treatment with any other AAV2-based gene therapy despite indication or are being considered for treatment with any other AAV2-based gene therapy? ☐ Yes ☐ No

Non-muscle invasive bladder cancer (NMIBC) with carcinoma in situ (CIS):

Does the member have a diagnosis of NMIBC with CIS with or without papillary tumors? ☐ Yes ☐ No

Is the member's disease high-risk and BCG unresponsive? ☐ Yes ☐ No

Is the member immunocompromised or immunodeficient? ☐ Yes ☐ No

Does the member have extra-vesical (i.e., urethra, ureter, or renal pelvis), muscle invasive (T2-T4), or metastatic urothelial carcinoma? ☐ Yes ☐ No

Is the member ineligible for or has elected not to undergo cystectomy? ☐ Yes ☐ No

Does the member have an ECOG performance status ≤2? ☐ Yes ☐ No

CURRENT or PREVIOUS THERAPY

Medication Name	Strength/ Frequency	Dates of Therapy	Status (Discontinued & Why/Current)

REAUTHORIZATION

Non-muscle invasive bladder cancer (NMIBC) with carcinoma in situ (CIS):

Does the member continue to meet the requirements for initial therapy and has been treated with nadofaragene firadenovec-vncg (Adstiladrin) with no adverse reactions? ☐ Yes ☐ No

Does the member have no signs of unacceptable toxicity (such as risk of disseminated adenovirus infection) while on treatment with nadofaragene firadenovec-vncg (Adstiladrin)? ☐ Yes ☐ No

SUPPORTING INFORMATION or CLINICAL RATIONALE

Prescribing Provider Signature	Date