SmartPA Criteria Proposal

Drug/Drug Class:	Enzyme Deficiency, Select Agents Clinical Edit
First Implementation Date:	October 20, 2022
Proposed Date:	October 17, 2023
Prepared for:	MO HealthNet
Prepared by:	MO HealthNet/Conduent
Criteria Status:	☐Existing Criteria ☒Revision of Existing Criteria ☐New Criteria

Executive Summary

Purpose: Ensure appropriate utilization and control of Enzyme Deficiency, Select Agents.

Why Issue Selected: Enzymes play an important role in the human body by carrying out various chemical functions such as digesting food, healing wounds, and breaking down toxins. Certain disease states result in deficiencies in enzymes and potentially lead to life-changing or life-threatening symptoms. Agents used to treat these disease states are sometimes classified as enzyme replacement therapies, which supplement the deficient enzyme, or have other mechanisms of action that result in increased enzyme levels in the body. By increasing levels of the deficient enzyme, these therapies treat the symptoms of the disease.

Due to the high cost and specific approved indications, MO HealthNet will impose clinical criteria to ensure appropriate utilization of Enzyme Deficiency, Select Agents.

Program-Specific Information:

Date Range FFS 7-1-22 to 6-30-23			
Drug	**Claims**	**Spend**	**Avg Spend per Claim**
ALDURAZYME 2.9 MG/5 ML VIAL	49	$1,525,863.20	$31,140.07
BRINEURA 150 MG/5 ML VIAL	1	$123,930.00	$123,930.00
CERDELGA 84 MG CAP	0	-	-
CEREZYME 400 UNIT VIAL	3	$52,023.90	$17,341.30
ELAPRASE 6 MG/3 ML VIAL	156	$2,386,062.56	$15,295.27
ELELYSO 200 UNIT VIAL	0	-	-
KANUMA 20 MG/10 ML VIAL	0	-	-
KUVAN 100 MG PWD PACKET	79	$190,854.00	$2,415.87
KUVAN 500 MG PWD PACKET	64	$679,447.70	$10,616.37
KUVAN 100 MG TAB	16	$140,418.96	$8,776.19
LAMZEDE 10 MG VIAL	0	-	-
MEPSEVII 10 MG/5 ML VIAL	0	-	-
MIGLUSTAT 100 MG CAPS	0	-	-
NAGLAZYME 1 MG/ML VIAL	0	-	-
NITISINONE 2 MG CAP	0	-	-
NITISINONE 5 MG CAP	0	-	-
NITISINONE 10 MG CAP	0	-	-
NITYR 2 MG TAB	0	-	-
NITYR 5 MG TAB	0	-	-

NITYR 10 MG TAB	6	$402,678.30	$67,113.05
ORFADIN 2 MG CAP	0	-	-
ORFADIN 5 MG CAP	0	-	-
ORFADIN 10 MG CAP	4	$218,024.45	$54,506.11
ORFADIN 20 MG CAP	0	-	-
ORFADIN 4 MG/ML SUSP	0	-	-
PALYNZIQ 2.5 MG/0.5 ML SYR	2	$3,587.04	$1,793.52
PALYNZIQ 10 MG/0.5 ML SYR	6	$31,045.96	$5,174.33
PALYNZIQ 20 MG/ML SYR	1	$17,855.55	$17,855.55
PYRUKYND 5 MG 28-DAY PACK	0	-	-
PYRUKYND 20 MG 28-DAY PACK	0	-	-
PYRUKYND 50 MG 28-DAY PACK	0	-	-
PRYUKYND 5 MG TAPER PACK	0	-	-
PYRUKYND 20/5 MG TAPER PACK	0	-	-
PYRUKYND 50/20 MG TAPER PACK	0	-	-
REVCOVI 2.4 MG/1.5 ML VIAL	0	-	-
RYPLAZIM 68.8 MG PWD FOR INJ	14	$516,097.35	$36,864.10
SAPROPTERIN 100 MG PWD PACKET	83	$155,336.95	$1,871.53
SAPROPTERIN 500 MG PWD PACK	119	$961,115.80	$8,076.60
SAPROPTERIN 100 MG TAB	25	$246,102.15	$9,844.09
STRENSIQ 18 MG/0.45 ML VIAL	15	$258,380.91	$17,225.39
STRENSIQ 28 MG/0.7 ML VIAL	28	$658,409.01	$23,514.61
STRENSIQ 40 MG/ML VIAL	30	$1,490,774.25	$49,692.48
STRENSIQ 80 MG/0.8 ML VIAL	58	$7,557,053.15	$130,294.02
VIMIZIM 1 MG/ML VIAL	0	-	-
VPRIV 400 UNIT VIAL	22	$329,654.81	$14,984.31
XENPOZYME 20 MG VIAL	0	-	-
XENPOZYME 4 MG VIAL	0	-	-
ZAVESCA 100 MG CAP	1	$26,828.85	$26,828.85

Type of Criteria: ☐ **Increased risk of ADE** ☐ **Preferred Drug List**
☒ **Appropriate Indications** ☒ **Clinical Edit**

Data Sources: ☐ **Only Administrative Databases** ☒ **Databases + Prescriber-Supplied**

Setting & Population

- Drug class for review: Enzyme Deficiency, Select Agents
- Age range: All appropriate MO HealthNet participants

Approval Criteria

Initial Therapy:

- Prescribed by or in consultation with an appropriate specialist in the treated disease state **AND**

- Documented diagnosis of acid sphingomyelinase deficiency (NPD-A/B or NPD-B):
  - Claim is for Xenpozyme AND
  - Diagnosis is confirmed by:

        - Molecular genetic testing confirming biallelic pathogenic variants in *SMDP1* gene OR
        - Documentation of deficient acid sphingomyelinase activity in peripheral leukocytes, cultured fibroblasts, or lymphocytes AND
    - Participants has clinical manifestations of ASMD, defined by ONE of the following:
        - Baseline DLco ≤ 70% of predicted normal OR
        - Spleen volume ≥ 6 MN for adults or ≥ 5 MN for participants < 18 years of age OR
        - Height ≤ -1 Z score, AND
    - Initial approval for 12 months

- Documented diagnosis of adenosine deaminase severe combined immune deficiency (ADA-SCID):
    - Claim is for Revcovi **AND**
    - Initial approval for 12 months

- **Documented diagnosis of alpha-mannosidosis (AM):**
    - **Claim is for Lamzede AND**
    - **Diagnosis confirmed by:**
        - **Molecular genetic testing revealing pathogenic variants of the MAN2B1 gene OR**
        - **Documentation of deficient acid alpha-mannosidase activity in leukocytes or other nucleated cells AND**
    - **Participant is aged 3 years or older AND**
    - **Participant is currently not pregnant**
    - **Initial approval for 12 months**

- Documented diagnosis of Gaucher disease:
    - Claim is for Cerdelga, Cerezyme, Elelyso, Vpriv, or Zavesca **AND**
    - For Cerdelga and Zavesca: participant is aged at least 18 years
    - Initial approval for 12 months

- Documented diagnosis of hereditary tyrosinemia type 1:
    - Claim is for Orfadin or Nityr **AND**
    - Initial approval for 12 months
    - For Orfadin 20 mg capsule: documentation of reason why participant cannot utilize lower strength capsules
    - For Nityr: documentation of reason why participant cannot utilize Orfadin

- Documented diagnosis of perinatal/infantile- and juvenile-onset hypophosphatasia (HPP):
    - Diagnosis confirmed by:
        - Presence of a known pathogenic variant in the *ALPL* gene as detected by *ALPL* molecular genetic testing **OR**
        - Diagnosis supported by all the following:
            - Radiographic imaging demonstrating skeletal abnormalities **AND**
            - Serum alkaline phosphatase (ALP) level below the gender- and age-specific reference range **AND**
            - Elevated tissue-nonspecific alkaline phosphatase substrate level **AND**
    - Disease onset prior to age 18 years **AND**
    - Participant has clinical manifestations of hypophosphatasia (i.e. skeletal abnormalities, respiratory problems, failure to thrive, rickets, etc.) **AND**
    - Claim is for Strensiq **AND**
    - Initial approval for 6 months

- Documented diagnosis of late-infantile neuronal ceroid lipofuscinosis type 2 (CLN2), tripeptidyl peptidase 1 (TPP1) deficiency:
    - Diagnosis confirmed by:
        - Deficient TPP1 enzyme activity in leukocytes, fibroblasts, or dried blood spots **OR**
        - Genetic testing confirming two pathogenic variants in the *TPP1*/*CLN2* gene **AND**

- Claim is for Brineura **AND**
- Participant has mild to moderate disease documented by a two-domain score of 3 to 6 on motor and language domains in the Hamburg CLN2 Clinical Rating Scale, with a score of at least 1 in each of these two domains **AND**
- Participant is ambulatory **AND**
- Documentation of baseline Hamburg CLN2 Clinical Rating Scale score
- Initial approval for 12 months

- Documented diagnosis of lysosomal acid lipase deficiency:
  - Claim is for Kanuma **AND**
  - Initial approval for 12 months

- Documented diagnosis of mucopolysaccharidosis I (MPS I):
  - Claim is for Aldurazyme **AND**
  - Initial approval for 12 months

- Documented diagnosis of mucopolysaccharidosis II (MPS II):
  - Claim is for Elaprase **AND**
  - Initial approval for 12 months

- Documented diagnosis of mucopolysaccharidosis IVA (MPS IVA):
  - Claim is for Vimizim **AND**
  - Initial approval for 12 months

- Documented diagnosis of mucopolysaccharidosis VI (MPS VI):
  - Claim is for Naglazyme **AND**
  - Initial approval for 12 months

- Documented diagnosis of mucopolysaccharidosis VII (MPS VII):
  - Claim is for Mepsevii **AND**
  - Initial approval for 12 months

- Documented diagnosis of phenylketonuria:
  - Claim is for Kuvan **AND**
  - **For Palynziq:**
    - **Documented adequate therapeutic trial of Kuvan (sapropterin) defined as 180 days of therapy in the past year or ADE/ADRs to Kuvan therapy AND**
    - **Documented baseline blood phenylalanine concentrations > 600 micromol/L**
  - Initial approval for 12 months

- Documented diagnosis of plasminogen deficiency type 1:
  - Diagnosis confirmed by:
    - Baseline plasminogen activity level ≤ 45% **AND**
    - Documented history of lesions (external and/or internal) and symptoms consistent with a diagnosis of plasminogen deficiency type 1 **AND**
    - Genetic testing confirming pathogenic variant in *PLG* gene **AND**
  - Claim is for Ryplazim **AND**
  - Initial approval for 12 months

- Documented diagnosis of symptomatic pyruvate kinase deficiency:
  - Diagnosis confirmed by:
    - Documentation of genetic testing confirming presence of at least 2 variant alleles in the *PKLR* gene, of which at least 1 is a missense variant **AND**
  - Documentation of previous red blood cell transfusions for hemolytic anemia in the past year **AND**

- Baseline hemoglobin level of ≤ 10 g/dL **AND**
- Claim is for Pyrukynd **AND**
- Initial approval for 6 months

- ~~**Documented diagnosis of urea cycle disorder:**~~
  - ~~**Diagnosis confirmed by enzymatic, biochemical, or genetic testing AND**~~
  - ~~**Claim is for Buphenyl, Pheburane Pellet, or Ravicti AND**~~
  - ~~**Documentation of trial and failure of dietary protein restriction and/or amino acid supplementation AND**~~
  - ~~**Initial approval for 12 months**~~
  - ~~**For Ravicti:**~~
    - ~~**Failure to achieve therapeutic response after minimum of 90 days of therapy with Buphenyl OR**~~
    - ~~**Documented ADE/ADR to Buphenyl**~~
  - ~~**For Pheburane: Clinical consultant review for medical necessity and reason why Buphenyl or Ravicti cannot be utilized**~~

Continuation of Therapy:

- Compliance to prescribed drug therapy **AND**
- For Brineura:
  - Documentation of benefit of therapy demonstrated by stabilization or lack of decline in motor function based on the Motor domain of the Hamburg CLN2 Clinical Rating Scale (decline is defined has having an unreversed (sustained) 2-category decline or an unreversed score of 0 in the Motor domain of the CLN2 Clinical Rating Scale)
  - Continued approval for 12 months
- For Pyrukynd:
  - Documentation of increase in hemoglobin of at least 1 g/dL from baseline **OR**
  - Documentation of reduction in transfusion burden from baseline
  - Continued approval for 12 months
- For Strensiq:
  - Documented benefit from therapy including one of the following:
    - Improved respiratory status
    - Improved growth from baseline
    - Improvement of skeletal manifestations from baseline
    - Lack of evidence of disease progression
  - Continued approval for 12 months
- For Xenpozyme:
  - Documented benefit from therapy including one of the following:
    - Improved height Z score from baseline
    - Reduced spleen volume from baseline
    - Improved DLco score from baseline
  - Continued approval for 12 months
- **For Lamzede:**
  - **Documentation of clinical benefit of therapy, such as slowed decline in the severity of signs and symptoms**
  - **Continued approval for 12 months**
- **For Palynziq:**
  - **Renewal of prior authorization may be given following documentation of blood phenylalanine concentrations < 600 micromol/L or at least 20% less than baseline level**
  - **Continued approval for 12 months**

Denial Criteria

- Therapy will be denied if all approval criteria are not met

- For Brineura:
  - Participant has acute intraventricular access device-related complication
  - Participant has ventriculoperitoneal shunts
- ~~**For Buphenyl/Ravicti/Pheburane Pellet: medication is being used for the treatment of acute hyperammonemia**~~
- **For Lamzede:**
  - **Paticipant has a history of haematopoietic stem cell transplantation (HSCT) or bone marrow transplant**
  - **Participant cannot walk without support**
- For Pyrukynd:
  - Participant is currently pregnant
  - Documentation of moderate to severe hepatic disease
  - Claim exceeds 2 tablets per day
- **For Palynziq:**
  - **Lack of an adequate response to therapy after 16 weeks of continuous treatment at the maximum dose of 60 mg daily**
  - **Participant is currently pregnant**
  - **Claim quantity exceeds 3 syringes per day**
- ~~**For Ravicti: documentation of N-acetylglutamate synthase (NAGS) deficiency**~~
- For Ryplazim: participant is currently pregnant
- For Xenpozyme:
  - Participant is currently pregnant
  - History of major organ transplant
  - Acute or rapidly progressive neurologic abnormalities

Required Documentation

Laboratory Results:		Progress Notes:	
MedWatch Form:		Other:	X

Disposition of Edit

Denial: Exception code “0682” (Clinical Edit)
Rule Type: CE

Default Approval Period

1 year

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