

Reference Number: LA.CP.CG.13 Date of Last Revision 01/25 Coding implications Revision Log

See <u>Important Reminder</u> at the end of this policy for important regulatory and legal information.

OVERVIEW

Hereditary metabolic disorders, also known as inborn errors of metabolism, are genetic disorders that interfere with the body's metabolism. There are hundreds of inherited metabolic disorders, and many are screened for at birth through newborn screening programs, while others are identified after a child or adult shows symptoms of the disorder. Genetic testing for metabolic disorders aids in quickly identifying the specific disorder so that proper treatment can be initiated and at-risk family members can be identified.

Hereditary endocrine disorders are a group of conditions involving the endocrine system, a network of glands that produce and release hormones in order to regulate body functions. This document aims to address hereditary endocrine disorders that are non-cancerous in nature.

MTHFR variants can affect metabolism of homocysteine; however, the treatment for elevated homocysteine levels is the same regardless of *MTHFR* status. The test is generally recognized as having little clinical utility and therefore is not generally recommended.

Mitochondrial disorders are a clinically heterogeneous group of disorders caused by dysfunction of the mitochondrial respiratory chain. The diagnosis of a primary mitochondrial disease can be difficult, as the individual symptoms are nonspecific and symptom patterns often overlap significantly. Mitochondrial disorders can be caused by mutations in the genes encoded by the mitochondrial DNA (mtDNA), which are transmitted by maternal inheritance, or by genes encoded by the nuclear DNA, which can be transmitted in an autosomal recessive or autosomal dominant manner. There are over 1000 nuclear genes coding for proteins that support mitochondrial function. These disorders can present at any age and many involve multiple organ systems, often with neurologic and myopathic features. Of note, a family history in which affected women transmit the disease to male and female children and affected men do not transmit the disease to their children suggests the familial variant(s) is in the mtDNA, rather than in a nuclear gene.

Genetic testing for metabolic, endocrine, and mitochondrial disorders aids in identifying the specific disorder that is present, so that proper treatment (if any) can be initiated, and at-risk family members can be identified.



POLICY REFERENCE TABLE

Coding Implications

This clinical policy references Current Procedural Terminology (CPT[®]). CPT is a registered trademark of the American Medical Association. All CPT codes and descriptions are copyrighted 2023, American Medical Association. All rights reserved. CPT codes and CPT descriptions are from the current manuals and those included herein are not intended to be all-inclusive and are included for informational purposes only. Codes referenced in this clinical policy are for informational purposes only and may not support medical necessity. Inclusion or exclusion of any codes does not guarantee coverage. Providers should reference the most up-to-date sources of professional coding guidance prior to the submission of claims for reimbursement of covered services.

NOTE: Coverage is subject to each requested code's inclusion on the corresponding LDH fee schedule. Non-covered codes are denoted (*) and are reviewed for Medical Necessity for members under 21 years of age on a per case basis. The non-covered codes will only be denoted in the table below and not throughout the policy. Please only reference the policy reference table for covered and non-covered codes.

The tests, associated laboratories, CPT codes, and ICD codes contained within this document serve only as examples to help users navigate claims and corresponding criteria; as such, they are not comprehensive and are not a guarantee of coverage or non-coverage. Please see the <u>Concert</u> <u>Platform</u> for a comprehensive list of registered tests.

Criteria Sections	Example Tests (Labs)	Common CPT Codes	Common ICD Codes	<u>Ref</u>		
MTHFR Variant Analysis						
<u>MTHFR Variant</u> Analysis	Methylenetetrahydrofolate Reductase (MTHFR) Thermolabile Variant, DNA Analysis (LabCorp)	81291*	E03.9, E55.9, E72.12, E78.2, E78.5, E88.9,	1, 2		
	Methylenetetrahydrofolate Reductase (MTHFR), DNA Mutation Analysis (Quest Diagnostics)		003, N96, R53.83, Z00.00			
Monogenic Diabetes (Including Maturity Onset Diabetes of the Young (MODY)						
Monogenic Diabetes (Including Maturity- Onset Diabetes of the	Maturity Onset Diabetes of the Young (MODY) Panel (PreventionGenetics, part of Exact Sciences)	81403*, 81405*, 81406*, 81407*, 81479	E10, E11, E16.1 E16.2	5, 11, 12		



Young (MODY)) Panels	Maturity-onset diabetes of the young (MODY) (Ambry Genetics)					
	Gene Evaluation (GCK,HNF1A,HNF1B,HNF4A,IPF1) (Athena Diagnostics Inc)					
Mitochondrial Genom	e Sequencing, Deletion/Duplication	, and/or Nuclear G	<u>enes</u>			
Mitochondrial Genome Sequencing, Deletion/Duplication, and/or Nuclear Gene Panel	Mito Genome Sequencing & Deletion Testing (GeneDx)	81460, 81465*	E88.40, E88.41, E88.42, E88.49,	3, 4		
	Mitochondrial Full Genome Analysis, Next-Generation Sequencing (NGS), Varies (Mayo Clinic Laboratories)		G31.82, H49.811- H49.819			
	Nuclear Mitochondrial Gene Panel, Next-Generation Sequencing, Varies (Mayo Clinic Laboratories)	81440*				
	MitoXpanded Panel (GeneDx)					
	Genomic Unity Comprehensive Mitochondrial Disorders Analysis (Variantyx)	0417U*				
Other Covered Metabo	Other Covered Metabolic, Endocrine, and Mitochondrial Disorders					
Other Covered Metabolic, Endocrine, and Mitochondrial Disorders	See list below	81400*-81408*, 81205*, 81250*		6, 7, 8, 9, 10		

OTHER RELATED POLICIES

This policy document provides criteria for metabolic, endocrine, and mitochondrial disorders. Please refer to:

- *Genetic Testing: Prenatal and Preconception Carrier Screening* for criteria related to prenatal or preconception **carrier** screening.
- *Genetic Testing: Prenatal Diagnosis (via amniocentesis, CVS, or PUBS) and Pregnancy Loss* for criteria related to prenatal and pregnancy loss **diagnostic** genetic testing.
- *Genetic Testing: Preimplantation Genetic Testing* for criteria related to genetic testing of embryos prior to in vitro fertilization.



- *Genetic Testing: Multisystem Inherited Disorders, Intellectual Disability, and Developmental Delay* for criteria related to genetic disorders that affect multiple organ systems.
- *Genetic Testing: Hereditary Cancer Susceptibility Syndromes* for criteria related to genetic testing for hereditary endocrine cancer predisposition syndromes.
- *Genetic Testing: General Approach to Genetic and Molecular Testing* for criteria related to metabolic, endocrine, and mitochondrial disorders not specifically discussed in this or another non-general policy, including known familial variant testing.

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CRITERIA

It is the policy of Louisiana Healthcare Connections that the specific genetic testing noted below is **medically necessary** when meeting the related criteria:

MTHFR VARIANT ANALYSIS

- I. *MTHFR* targeted variant analysis (e.g., 677T, 1298C) (81291) is considered **investigational** for all indications, including but not limited to:
 - A. Evaluation for thrombophilia or recurrent pregnancy loss
 - B. Evaluation of at-risk relatives
 - C. Drug metabolism, such as in pharmacogenetic testing.

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MONOGENIC DIABETES (INCLUDING MATURITY-ONSET DIABETES OF THE YOUNG (MODY))

Monogenic Diabetes (Including Maturity Onset Diabetes of the Young (MODY)) Panels

- I. Multigene panel analysis to establish or confirm a diagnosis of monogenic diabetes (including maturity-onset diabetes of the young (MODY)) (81403, 81405, 81406, 81407, 81479) is considered **medically necessary** when:
 - A. The member/enrollee has a diagnosis of diabetes within the first 12 months of life, **OR**
 - B. The member/enrollee has a diagnosis of diabetes before 30 years of age, AND



- 1. The member/enrollee has at least one of the following:
 - a) Autoantibody negative, **OR**
 - b) Retained C-peptide levels, **OR**
- C. The member/enrollee has a diagnosis of diabetes not characteristic of type 1 or type 2 diabetes, **AND**
 - 1. The member/enrollee has a family history of diabetes consistent with an <u>autosomal dominant pattern of inheritance</u>.
- II. Multigene panel analysis to establish or confirm a diagnosis of monogenic diabetes (maturity-onset diabetes of the young (MODY)) (81403, 81404, 81405, 81406, 81407, 81479) is considered **investigational** for all other indications.

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MITOCHONDRIAL GENOME SEQUENCING, DELETION/DUPLICATION, AND/OR NUCLEAR GENES

Mitochondrial Genome Sequencing, Deletion/Duplication, and/or Nuclear Genes

- I. Mitochondrial genome sequencing (81460), deletion/duplication (81465), and/or nuclear genes analysis (0417U, 81440) to establish or confirm a diagnosis of a primary mitochondrial disorder is considered **medically necessary** when:
 - A. The member/enrollee has a classic phenotype of one of the maternally inherited syndromes (e.g., <u>Leber hereditary optic neuropathy</u>, <u>mitochondrial</u> <u>encephalomyopathy with lactic acidosis and stroke-like episodes [MELAS]</u>, <u>myoclonic epilepsy with ragged red fibers [MERRF]</u>, maternally inherited deafness and diabetes [MIDD], neuropathy, ataxia, retinitis pigmentosa [NARP], Kearns-Sayre syndrome/CPEO); or of a nuclear DNA mitochondrial disorder (e.g., <u>mitochondrial neurogastrointestinal encephalopathy [MNGIE]</u>); **OR**
 - B. The member/enrollee has non-specific clinical features suggestive of a primary mitochondrial disorder and meets **ALL** of the following:
 - 1. Clinical findings of at least two of the following:
 - a) Ptosis, OR
 - b) External ophthalmoplegia, **OR**
 - c) Proximal myopathy, OR
 - d) Exercise intolerance, OR



- e) Cardiomyopathy, **OR**
- f) Sensorineural deafness, **OR**
- g) Optic atrophy, **OR**
- h) Pigmentary retinopathy, **OR**
- i) Diabetes mellitus, **OR**
- j) Fluctuating encephalopathy, **OR**
- k) Seizures, OR
- l) Dementia, OR
- m) Migraine, OR
- n) Stroke-like episodes, OR
- o) Ataxia, **OR**
- p) Spasticity, OR
- q) Chorea, **OR**
- r) Multiple late term pregnancy loss, AND
- 2. Conventional biochemical laboratory studies have been completed and are non-diagnostic, including at least: plasma or CSF lactic acid concentration, ketone bodies, plasma acylcarnitines, and urinary organic acids, **AND**
- 3. Additional diagnostic testing indicated by the member/enrollee's clinical presentation (e.g., fasting blood glucose, electrocardiography, neuroimaging, electromyography, echocardiography, audiology, thyroid testing, electroencephalography, exercise testing) have been completed and are non-diagnostic.
- II. Mitochondrial genome sequencing (81460), deletion/duplication (81465), and/or nuclear genes analysis (81440) to establish or confirm a diagnosis of a primary mitochondrial disorder is considered **investigational** for all other indications.

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OTHER COVERED METABOLIC, ENDOCRINE, AND MITOCHONDRIAL DISORDERS

The following is a list of conditions that have a known genetic association. Due to their relative rareness, it may be appropriate to cover these genetic tests to establish or confirm a diagnosis.

- I. Genetic testing to establish or confirm one of the following metabolic, endocrine, and mitochondrial conditions to guide management is considered **medically necessary** when the member/enrollee demonstrates clinical features* consistent with the disorder (the list is not meant to be comprehensive, see II below):
 - A. Congenital adrenal hyperplasia, including:
 - 1. <u>21-Hydroxylase deficiency</u>



- B. Congenital disorders of glycosylation
- C. Congenital hyperinsulinism
- D. Disorders of amino acid and peptide metabolism, including:
 - 1. <u>Glutaric acidemia type I (GA-1)</u>
 - 2. Homocystinuria caused by cystathionine beta-synthase (CBS) deficiency
 - 3. <u>Methylmalonic acidemia</u>
 - 4. Propionic acidemia
 - 5. <u>Maple Syrup Urine Disease (MSUD)</u>
- E. Disorders of biotin metabolism, including:
 - 1. Biotinidase deficiency
- F. Disorders of carnitine transport and the carnitine cycle, including:
 - 1. Carnitine palmitoyltransferase II deficiency
 - 2. Primary carnitine deficiency
- G. Disorders of copper metabolism, including:
 - 1. <u>ATP7A-Related copper transport disorders</u> (e.g., Menkes disease, occipital horn syndrome (OHS), ATP7A-related distal motor neuropathies)
 - 2. Wilson disease
- H. Disorders of fatty acid oxidation, including:
 - 1. <u>Medium-chain acyl-coenzyme A dehydrogenase deficiency (MCAD deficiency)</u>
- I. Disorders of galactose metabolism, including:
 - 1. Galactosemia
- J. Disorders of glucose transport, including:
 - 1. <u>Glucose transporter type I deficiency syndrome (Glut1 DS)</u>
- K. Disorders of phenylalanine or tyrosine metabolism, including:
 - 1. <u>Alkaptonuria</u>
 - 2. <u>Phenylalanine hydroxylase deficiency</u>
- L. Disorders of porphyrin and heme metabolism, including:
 - 1. Acute intermittent porphyria
- M. Fibrous Dysplasia/McCune-Albright Syndrome
- N. Glycogen storage disorders, including:
 - 1. <u>Glycogen Storage Disease Type I (GSDI)</u>
 - 2. <u>Pompe disease (GSDII)</u>
- O. <u>Hypophosphatasia</u>
- P. Kallmann syndrome (GnRH deficiency)
- Q. Lysosomal storage disorders, including:
 - 1. Gaucher disease
 - 2. <u>Krabbe disease</u>
 - 3. <u>MPS-Type I (Hurler syndrome)</u>
 - 4. <u>MPS-Type II (Hunter syndrome)</u>
 - 5. <u>Mucolipidosis IV</u>
- R. Urea cycle disorders, including:
 - 1. <u>Ornithine Transcarbamylase (OTC) deficiency</u>
- S. Malignant hyperthermia



T. SHOX deficiency disorders.

II. Genetic testing to establish or confirm the diagnosis of all other metabolic, endocrine, and <u>mitochondrial disorders</u> not specifically discussed within this or another medical policy will be evaluated by the criteria outlined in *General Approach to Genetic and Molecular Testing* (see policy for criteria).

*Clinical features for a specific disorder may be outlined in resources such as <u>GeneReviews</u>, <u>OMIM</u>, <u>National Library</u> <u>of Medicine</u>, <u>Genetics Home Reference</u>, or other scholarly sources.

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DEFINITIONS

- 1. **Mitochondrial disease** refers to a heterogenous group of disorders caused by dysfunctional mitochondria, the organelles responsible for oxidative phosphorylation within the cell.
- 2. Autosomal dominant inheritance refers to a type of transmission of a genetic condition in which only one mutated copy of a gene (rather than two) is necessary for an individual to manifest the disease. The mutation can be inherited from either parent, and the disease can typically be seen in any sex. A pedigree (family history) that has an autosomal dominant disorder will typically have affected family members in each generation, though some family members may be more severely affected than others.

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BACKGROUND AND RATIONALE

METHYLENETETRAHYDROFOLATE REDUCTASE (MTHFR)

MTHFR Variant Analysis

American College of Medical Genetics and Genomics (ACMG)

ACMG published a practice guideline for *MTHFR* polymorphism testing (2013, confirmed 2020) with the following recommendations:

- *MTHFR* polymorphism genotyping should not be ordered as part of the clinical evaluation for thrombophilia or recurrent pregnancy loss
- *MTHFR* polymorphism genotyping should not be ordered for at-risk family members (p. 154)



MONOGENIC DIABETES (INCLUDING MATURITY-ONSET DIABETES OF THE YOUNG (MODY))

Monogenic Diabetes (Including Maturity-Onset Diabetes of the Young (MODY)) Panels

American Diabetes Association

In 2021, the American Diabetes Association made the following recommendations:

- All children diagnosed with diabetes in the first 6 months of life should have immediate genetic testing for neonatal diabetes. (Category A)
- Children and those diagnosed in early adulthood who have diabetes not characteristic of type 1 or type 2 diabetes that occurs in successive generations (suggestive of an autosomal dominant pattern of inheritance) should have genetic testing for maturity-onset diabetes of the young. (Category A)
- In both instances, consultation with a center specializing in diabetes genetics is recommended to understand the significance of these mutations and how best to approach further evaluation, treatment, and genetic counseling. (Category E) (p. 525)

Murphy, et al.

Murphy, et al (2023) performed a systematic review and issued an expert opinion on how to use precision diagnostics to identify individuals with monogenic diabetes. The article states that the following individuals should be offered testing for monogenic diabetes:

- 1. All patients diagnosed with diabetes before the age of 6 months should be tested for monogenic forms of neonatal diabetes using the large-gene panel.
- 2. All patients diagnosed between 6 and 12 months should be tested for monogenic forms of neonatal diabetes using the large-gene panel. No demonstrable yield of monogenic etiology to support reflexive genetic testing patients diagnosed with diabetes between 12-24 months.
- 3. Women with gestational diabetes and fasting glucose above 5.5 mmol/L without obesity* should be tested for GCK etiology.
- 4. Those with persisting, mild hyperglycemia (HbA1c 38–62 mmol/mol, or fasting glucose 5.5–7.8 mmol/L) at any age, in the absence of obesity* should be tested for GCK etiology.
- 5. People without obesity under the age of 30 years who are either autoantibody negative and/or have retained C-peptide levels should be tested for monogenic diabetes using a large-gene panel. (p.10)

International Society for Pediatric and Adolescent Diabetes (ISPAD)

In 2022, the International Society for Pediatric and Adolescent Diabetes (ISPAD) released a clinical practice consensus guideline for the diagnosis and management of monogenic diabetes in children and adolescents. The statement includes the following recommendations for genetic testing in the setting of neonatal diabetes and maturity onset diabetes of the young:



"All infants diagnosed with diabetes in the first 6 months of life are recommended to have immediate molecular genetic testing. Genetic testing may be considered in infants diagnosed between 6 and 12 months, especially in those without islet autoantibodies or who have other features suggestive of a monogenic cause." (p. 1190)

"The diagnosis of maturity onset diabetes of the young (MODY) is recommended in the following scenarios: family history of diabetes in a parent and first-degree relatives of that affected parent in persons with diabetes who lack the characteristics of T1D and T2D." (p. 1191)

Mitochondrial Genome Sequencing, Deletion/Duplication, and/or Nuclear Genes

Mitochondrial Medicine Society

The Mitochondrial Medicine Society (2015) published the following consensus recommendations for DNA testing for mitochondrial disorders:

- 1. Massively parallel sequencing/NGS of the mtDNA genome is the preferred methodology when testing mtDNA and should be performed in cases of suspected mitochondrial disease instead of testing for a limited number of pathogenic point mutations.
- 2. Patients with a strong likelihood of mitochondrial disease because of a mtDNA mutation and negative testing in blood, should have mtDNA assessed in another tissue to avoid the possibility of missing tissue-specific mutations or low levels of heteroplasmy in blood; tissue-based testing also helps assess the risk of other organ involvement and heterogeneity in family members and to guide genetic counseling.
- 3. Heteroplasmy analysis in urine can selectively be more informative and accurate than testing in blood alone, especially in cases of MELAS due to the common m. 3243A>G mutation.
- 4. mtDNA deletion and duplication testing should be performed in cases of suspected mitochondrial disease via NGS of the mtDNA genome, especially in all patients undergoing a diagnostic tissue biopsy.
 - a. If a single small deletion is identified using polymerase chain reaction-based analysis, then one should be cautious in associating these findings with a primary mitochondrial disorder.
 - b. When multiple mtDNA deletions are noted, sequencing of nuclear genes involved in mtDNA biosynthesis is recommended.
- 5. When a tissue specimen is obtained for mitochondrial studies, mtDNA content (copy number) testing via real-time quantitative polymerase chain reaction should strongly be considered for mtDNA depletion analysis because mtDNA depletion may not be detected in blood.



- a. mtDNA proliferation is a nonspecific compensatory finding that can be seen in primary mitochondrial disease, secondary mitochondrial dysfunction, myopathy, hypotonia, and as a by-product of regular, intense exercise.
- 6. When considering nuclear gene testing in patients with likely primary mitochondrial disease, NGS methodologies providing complete coverage of known mitochondrial disease genes is preferred. Single-gene testing should usually be avoided because mutations in different genes can produce the same phenotype. If no known mutation is identified via known NGS gene panels, then whole exome sequencing should be considered. (p. 692-693)

GeneReviews: Primary Mitochondrial Disorders Overview

GeneReviews is an expert-authored review of current literature on a genetic disease, and goes through a rigorous editing and peer review process before being published online. Their recommendations are as follows:

Common clinical features of mitochondrial disorders include:

- ptosis
- external ophthalmoplegia
- proximal myopathy
- exercise intolerance
- cardiomyopathy
- sensorineural deafness
- optic atrophy
- pigmentary retinopathy
- diabetes mellitus
- fluctuating encephalopathy
- seizures
- dementia
- migraine
- stroke-like episodes
- ataxia
- spasticity
- chorea
- high incidence of mid- and late-pregnancy loss

When a patient's clinical picture is nonspecific but highly suggestive of a mitochondrial disorder, the clinician should start with measurement of plasma or CSF lactic acid concentration, ketone bodies, plasma acylcarnitines, and urinary organic acids.

Traditionally, the diagnosis of mitochondrial disorders has been based on demonstrating mitochondrial dysfunction in a relevant tissue biopsy (e.g., a skeletal muscle or liver biopsy, or skin fibroblasts), with the particular pattern of biochemical abnormality being used to direct targeted molecular genetic testing of mtDNA, specific nuclear genes, or both.



However, the more widespread availability of molecular diagnostic techniques and the advent of exome and genome sequencing has changed the diagnostic approach.

One important caveat arises from the fact that many mtDNA pathogenic variants are heteroplasmic, and the proportion of mutated mtDNA in blood may be undetectable. This can be circumvented by analyzing mtDNA from another tissue – typically skeletal muscle or urinary epithelium – in which the level of heteroplasmy tends to be higher. Some common mtDNA pathogenic variants (e.g., large-scale deletions causing CPEO) may only be detected in skeletal muscle.

In individuals with a specific clinical phenotype (e.g., MELAS, LHON, POLG-related disorders) it may be possible to reach a diagnosis with targeted analysis of specific mtDNA pathogenic variants or single-gene testing of a nuclear gene.

A mitochondrial disorders multigene panel is most likely to identify the genetic cause of the condition while limiting identification of variants of uncertain significance and pathogenic variants in genes that do not explain the underlying phenotype.

Comprehensive genomic testing does not require the clinician to determine which gene is likely involved. Such testing includes exome sequencing, genome sequencing, and mitochondrial sequencing which can simultaneously analyze nuclear DNA and mtDNA.

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Reviews, Revisions, and Approvals		Approval Date	Effective Date
Converted corporate to local policy.	09/23	11/27/23	
Semi-annual review. Updated title to reflect V1.2024 version. Overview, coding, reference-table, background and references updated. Throughout policy: replaced "coverage criteria" with "criteria. For MTHFR Variant Analysis: under I. added "but not limited to:". For Mitochondrial Genome Sequencing, Deletion/Duplication, and/or Nuclear Genes Panel: under I.B.1.i. removed "and deafness". For Other Covered Metabolic, Endocrine, and Mitochondrial Disorders Panel: under II. Added "and Molecular"; added II.3. "Autosomal dominant inheritance". For Background and Rationale; Known Familial Variant Analysis for Metabolic, Endocrine, and Mitochondrial Disorders panel: replaced "inheritance patterns" with "genetic testing"; under MTHFR Variant Analysis: added ". Confirmed 2020".		4/26/24	
Semi-annual review. Updated title to reflect V2.2024 version. In Known Familial Variant Analysis for Metabolic, Endocrine, and Mitochondrial Disorders, moved criteria to policy "Genetic Testing: General Approach to Genetic and Molecular Testing" to consolidate criteria for known familial variant tests. In Monogenic Diabetes (Including Maturity-Onset Diabetes of the Young (MODY)) Panels, criteria set name changed (formerly "Maturity- Onset Diabetes of the Young (MODY)"). Minor rewording for clarity throughout. Coding, reference-table, background and references updated.		9/17/24	10/17/24



Reviews, Revisions, and Approvals	Revision	Approval	Effective
	Date	Date	Date
Semi-annual review. Updated title to reflect V1.2025. MTHFR Variant Analysis: Streamlined portions of Background and Rationale section for brevity.	1/25	3/31/25	5/1/25

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Important Reminder

This clinical policy has been developed by appropriately experienced and licensed health care professionals based on a review and consideration of currently available generally accepted standards of medical practice; peer-reviewed medical literature; government agency/program approval status; evidence-based guidelines and positions of leading national health professional organizations; views of physicians practicing in relevant clinical areas affected by this clinical policy; and other available clinical information. LHCC makes no representations and accepts no liability with respect to the content of any external information used or relied upon in developing this clinical policy. This clinical policy is consistent with standards of medical practice current at the time that this clinical policy was approved.

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