Clinical Policy: DNA Analysis of Stool to Screen for Colorectal Cancer

Reference Number: WNC.CP.137
Last Review Date: 08/21

Coding Implications
Revision Log

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

Note: When state Medicaid coverage provisions conflict with the coverage provisions in this clinical policy, state Medicaid coverage provisions take precedence. Please refer to the state Medicaid manual for any coverage provisions pertaining to this clinical policy.

Description

Cologuard is a noninvasive screening test for colon cancer. This test comprises a multi-target screen for several aberrant DNA markers of colon cancer, as well as a hemoglobin immunoassay. This policy describes the medical necessity requirements for DNA analysis of stool with Cologuard.

Policy/Criteria

I. It is the policy of WellCare of North Carolina® that screening for colorectal cancer by DNA analysis of stool (i.e., Cologuard) is medically necessary **every year** when meeting the following:
   A. Age 45-85 years;
   B. Asymptomatic and at average risk for colon cancer;
   C. Is not within the standard interval of another screening test for colon cancer.

II. It is the policy of WellCare of North Carolina® that evidence based clinical and preventive guidelines do not support the use of DNA analysis of stool (i.e., Cologuard) for any circumstances other than those specified above.

Background

Colorectal cancer has become the second leading cause of cancer-related deaths in the United States, according to the latest statistics.[3] Multi-target stool testing for colorectal cancer is a noninvasive DNA test that screens for multiple lesions, including those related to Kras mutations, NDRG4 and BMP3 methylations, β-actin, and hemoglobin immunoassay.[1] The FDA approved Cologuard (Exact Sciences) based on this multi-target stool testing.[2] The sensitivity for detecting colorectal cancer from the multi-target DNA testing was 92.3% (60 of 65) and 73.8% (48 of 65) with fecal immunohistochemical tests (FIT), which look for intact human hemoglobin. Multi-target DNA testing is not a replacement for diagnostic colonoscopy testing in patients at high risk for colorectal cancer.

American Cancer Society

2018 Guidelines by the ACS give a qualified recommendation for screening for colorectal cancer starting at age 45. A qualified recommendation "indicates there is clear evidence of benefit of screening but less certainty about the balance of benefits and harms, or about patients' values and preferences, which could lead to different decisions about screening." The ACS gives a strong recommendation that colorectal cancer screening be performed in adults aged 50-75, and a qualified recommendation for adults aged 76-85.

United States Preventative Services Task Force (USPSTF)
The USPSTF recommends screening for colorectal cancer starting at age 50 years and continuing until age 75 years. According to this recommendation, the specificity of FIT-DNA is lower than that of FIT alone, and has a higher number of false-positive results, as well as a higher likelihood of follow-up colonoscopy and associated adverse events per screening test.[3] While no longitudinal follow-up data exists, with an abnormal FIT-DNA test result followed by a negative colonoscopy, there is potential for overly intensive surveillance due to clinician and patient concerns about the implications of the genetic component of the test.[3]
National Comprehensive Cancer Network (NCCN)
NCCN recommends the inclusion of multitarget stool DNA testing as a potential screening modality in average-risk individuals, but data to help determine an appropriate interval between screening, adherence to/participation rates of screening, and how multitarget stool DNA testing may fit into an overall screening program are limited, noting also that there are "no or limited data in high-risk individuals and the use of stool DNA should be individualized." NCCN recommends colorectal cancer screening for average-risk individuals 50-75 years of age, and on an individualized basis for those 76-85 years of age.
Multi-Society Task Force for Colorectal Cancer
The American College of Gastroenterology, the American Gastroenterological Association, and the American Society for Gastrointestinal Endoscopy issued a joint statement recommending FIT-fecal DNA tests every 3 years, as a second-tier screening tool for colorectal cancer. They offer a strong recommendation, based on high-quality evidence, for colorectal cancer screening beginning at age 50. Based on limited evidence and the high incidence of colorectal cancer in African-Americans, they recommend screening for this population starting at age 45.

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 2019, 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. 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.

CPT®* Codes	Description
81528	Oncology (colorectal) screening, quantitative real time target and signal amplification of 10 DNA markers (KRAS mutations, promoter methylation of NDRG4 and BMP3) and fecal hemoglobin, utilizing stool, algorithm reported as a positive or negative result

HCPCS ®* Codes	Description
No applicable codes.	

ICD-10-CM Diagnosis Codes that Support Coverage Criteria
+ Indicates a code(s) requiring an additional character

ICD-10-CM Code	Description
Z12.11	Encounter for screening for malignant neoplasm of colon
Z12.12	Encounter for screening for malignant neoplasm of rectum

Reviews, Revisions, and Approvals	Date	Approval Date
Original approval date	03/21	05/21
Revised criteria to yearly screening. Reviewed CPT and ICD-10-CM codes. Updated references.	08/21	

References

1. Imperiale TF, Ransohoff DF, Itzkowitz SH, et al. Multitarget stool DNA testing for colorectal-cancer screening. N Engl J Med 2014;370:1287-97.
2. Abramowicz, Mark, Gianna Zuccotti, and Jean-Marie Pflomm. A stool DNA test (Cologuard) for colorectal cancer screening." JAMA. 2014;312(23).
3. US Preventive Services Task Force. Final Recommendation Statement: Screening for Colorectal Cancer, May 18, 2021. Accessed August 24, 2021. https://uspreventiveservicestaskforce.org/uspstf/recommendation/colorectal-cancer-screening
4. National Comprehensive Cancer Network. NCCN clinical practice guidelines in oncology: Colorectal cancer screening. Version 2.2020. Accessed June 11, 2020.
5. Centers for Medicare and Medicaid Services. National Coverage Determination (NCD) for Colorectal Cancer Screening Tests (210.3). CMS.gov. Effective October 9, 2014. Accessed June 11, 2020.
6. American Cancer Society. American Cancer Society Guideline for Colorectal Cancer Screening: A Summary for Clinicians. American Cancer Society. 2018. Accessed June 11, 2020.
7. Siegel RL, Fedewa SA, Anderson WF, et al. Colorectal Cancer Incidence Patterns in the United States, 1974–2013. J Natl Cancer Inst. 2017 Aug 1;109(8). doi: 10.1093/jnci/djw322.
8. Rex DK, Boland CR, Dominitz JA, et al. Colorectal Cancer Screening: Recommendations for Physicians and Patients from the U.S. Multi-Society Task Force on Colorectal Cancer. Am J Gastroenterol. 2017;112(7):1016. Epub 2017 Jun 6.
9. Doubeni C. Screening for colorectal cancer: Strategies in patients at average risk. In: UpToDate. Lamont JT, Elmore JG (Eds). UpToDate, Waltham, MA. Accessed June 11, 2020
10. Doubeni C. Tests for screening for colorectal cancer In: UpToDate. Lamont JT, Elmore JG (Eds). March 18, 2020. Accessed June 11, 2020

North Carolina Guidance
Eligibility Requirements

a. An eligible beneficiary shall be enrolled in either:
   1. the NC Medicaid Program (Medicaid is NC Medicaid program, unless context clearly indicates otherwise); or
   2. the NC Health Choice (NCHC is NC Health Choice program, unless context clearly indicates otherwise) Program on the date of service and shall meet the criteria in this policy.

b. Provider(s) shall verify each Medicaid or NCHC beneficiary's eligibility each time a service is rendered.

c. The Medicaid beneficiary may have service restrictions due to their eligibility category that would make them ineligible for this service.

d. Following is only one of the eligibility and other requirements for participation in the NCHC Program under GS 108A-70.21(a): Children must be between the ages of 6 through 18.

EPSDT Special Provision: Exception to Policy Limitations for a Medicaid Beneficiary under 21 Years of Age

a. 42 U.S.C. § 1396d(r) [1905(r) of the Social Security Act]

Early and Periodic Screening, Diagnostic, and Treatment (EPSDT) is a federal Medicaid requirement that requires the state Medicaid agency to cover services, products, or procedures for Medicaid beneficiary under 21 years of age if the service is medically necessary health care to correct or ameliorate a defect, physical or mental illness, or a condition [health problem] identified through a screening examination (includes any evaluation by a physician or other licensed practitioner).

This means EPSDT covers most of the medical or remedial care a child needs to improve or maintain his or her health in the best condition possible, compensate for a health problem, prevent it from worsening, or prevent the development of additional health problems.

Medically necessary services will be provided in the most economic mode, as long as the treatment made available is similarly efficacious to the service requested by the beneficiary's physician, therapist, or other licensed practitioner; the determination process does not delay the delivery of the needed service; and the determination does not limit the beneficiary's right to a free choice of providers.

EPSDT does not require the state Medicaid agency to provide any service, product or procedure:

1. that is unsafe, ineffective, or experimental or investigational.
2. that is not medical in nature or not generally recognized as an accepted method of medical practice or treatment.

Service limitations on scope, amount, duration, frequency, location of service, and other specific criteria described in clinical coverage policies may be exceeded or may not apply as long as the provider's documentation shows that the requested service is medically necessary "to correct or ameliorate a defect, physical or mental illness, or a condition" [health problem]; that is, provider documentation shows how the service, product, or procedure meets all EPSDT criteria, including to correct or improve or maintain the beneficiary's health

in the best condition possible, compensate for a health problem, prevent it from worsening, or prevent the development of additional health problems.

EPSDT and Prior Approval Requirements

1. If the service, product, or procedure requires prior approval, the fact that the beneficiary is under 21 years of age does NOT eliminate the requirement for prior approval.
2. **IMPORTANT ADDITIONAL INFORMATION** about EPSDT and prior approval is found in the *NCTracks Provider Claims and Billing Assistance Guide*, and on the EPSDT provider page. The Web addresses are specified below:
   NCTracks Provider Claims and Billing Assistance Guide:
   https://www.nctracks.nc.gov/content/public/providers/provider-manuals.html
   EPSDT provider page: https://medicaid.ncdhhs.gov/

EPSDT does not apply to NCHC beneficiaries.

Provider(s) Eligible to Bill for the Procedure, Product, or Service
To be eligible to bill for the procedure, product, or service related to this policy, the provider(s) shall:

a. meet Medicaid or NCHC qualifications for participation;
b. have a current and signed Department of Health and Human Services (DHHS) Provider Administrative Participation Agreement; and
c. bill only for procedures, products, and services that are within the scope of their clinical practice, as defined by the appropriate licensing entity.

Compliance
Provider(s) shall comply with the following in effect at the time the service is rendered:

a. All applicable agreements, federal, state and local laws and regulations including the Health Insurance Portability and Accountability Act (HIPAA) and record retention requirements; and
b. All NC Medicaid's clinical (medical) coverage policies, guidelines, policies, provider manuals, implementation updates, and bulletins published by the Centers for Medicare and Medicaid Services (CMS), DHHS, DHHS division(s) or fiscal contractor(s).

Claims-Related Information
Provider(s) shall comply with the, NC Tracks Provider Claims and Billing Assistance Guide, Medicaid bulletins, fee schedules, NC Medicaid's clinical coverage policies and any other relevant documents for specific coverage and reimbursement for Medicaid and NCHC:

a. Claim Type - as applicable to the service provided:
   Professional (CMS-1500/837P transaction)
   Institutional (UB-04/837I transaction)
   Unless directed otherwise, Institutional Claims must be billed according to the National Uniform Billing Guidelines. All claims must comply with National Coding Guidelines.
b. International Classification of Diseases and Related Health Problems, Tenth Revisions, Clinical Modification (ICD-10-CM) and Procedural Coding System (PCS) - Provider(s) shall report the ICD-10-CM and Procedural Coding System (PCS) to the highest level of specificity that supports medical necessity. Provider(s) shall use the current ICD-10

edition and any subsequent editions in effect at the time of service. Provider(s) shall refer to the applicable edition for code description, as it is no longer documented in the policy.

c. Code(s) - Provider(s) shall report the most specific billing code that accurately and completely describes the procedure, product or service provided. Provider(s) shall use the Current Procedural Terminology (CPT), Health Care Procedure Coding System (HCPCS), and UB-04 Data Specifications Manual (for a complete listing of valid revenue codes) and any subsequent editions in effect at the time of service. Provider(s) shall refer to the applicable edition for the code description, as it is no longer documented in the policy. If no such specific CPT or HCPCS code exists, then the provider(s) shall report the procedure, product or service using the appropriate unlisted procedure or service code.
   Unlisted Procedure or Service
   CPT: The provider(s) shall refer to and comply with the Instructions for Use of the CPT Codebook, Unlisted Procedure or Service, and Special Report as documented in the current CPT in effect at the time of service.
   HCPCS: The provider(s) shall refer to and comply with the Instructions For Use of HCPCS National Level II codes, Unlisted Procedure or Service and Special Report as documented in the current HCPCS edition in effect at the time of service
d. Modifiers - Providers shall follow applicable modifier guidelines.
e. Billing Units - Provider(s) shall report the appropriate code(s) used which determines the billing unit(s).
f. Co-payments -
   For Medicaid refer to Medicaid State Plan:
   https://medicaid.ncdhhs.gov/get-involved/nc-health-choice-state-plan
   For NCHC refer to NCHC State Plan:
   https://medicaid.ncdhhs.gov/get-involved/nc-health-choice-state-plan
g. Reimbursement - Provider(s) shall bill their usual and customary charges. For a schedule of rates, refer to: https://medicaid.ncdhhs.gov/.

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. The Health Plan 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. “Health Plan” means a health plan that has adopted this clinical policy and that is operated or administered, in whole or in part, by Centene Management Company, LLC, or any of such health plan’s affiliates, as applicable.

The purpose of this clinical policy is to provide a guide to medical necessity, which is a component of the guidelines used to assist in making coverage decisions and administering benefits. It does not constitute a contract or guarantee regarding payment or results. Coverage

decisions and the administration of benefits are subject to all terms, conditions, exclusions and limitations of the coverage documents (e.g., evidence of coverage, certificate of coverage, policy, contract of insurance, etc.), as well as to state and federal requirements and applicable Health Plan-level administrative policies and procedures.

This clinical policy is effective as of the date determined by the Health Plan. The date of posting may not be the effective date of this clinical policy. This clinical policy may be subject to applicable legal and regulatory requirements relating to provider notification. If there is a discrepancy between the effective date of this clinical policy and any applicable legal or regulatory requirement, the requirements of law and regulation shall govern. The Health Plan retains the right to change, amend or withdraw this clinical policy, and additional clinical policies may be developed and adopted as needed, at any time.

This clinical policy does not constitute medical advice, medical treatment or medical care. It is not intended to dictate to providers how to practice medicine. Providers are expected to exercise professional medical judgment in providing the most appropriate care, and are solely responsible for the medical advice and treatment of members/enrollees. This clinical policy is not intended to recommend treatment for members/enrollees. Members/enrollees should consult with their treating physician in connection with diagnosis and treatment decisions.

Providers referred to in this clinical policy are independent contractors who exercise independent judgment and over whom the Health Plan has no control or right of control. Providers are not agents or employees of the Health Plan.

This clinical policy is the property of the Health Plan. Unauthorized copying, use, and distribution of this clinical policy or any information contained herein are strictly prohibited. Providers, members/enrollees and their representatives are bound to the terms and conditions expressed herein through the terms of their contracts. Where no such contract exists, providers, members/enrollees and their representatives agree to be bound by such terms and conditions by providing services to members/enrollees and/or submitting claims for payment for such services.