



Reimbursement Policy

Policy Number: RPLAB037

Policy Title: Serum Tumor Markers for Malignancies

Approval Date: May 15, 2026

Effective Date: Sept. 4, 2026

Policy Disclaimer

If a conflict arises between a Reimbursement Policy and any Plan document under which a member is entitled to covered services, the Plan document will govern. If a conflict arises between a reimbursement policy and any provider contract pursuant to which a provider participates in and/or provides covered services to eligible member(s) and/or plans, the provider's contract will govern. "Plan documents" include, but are not limited to, Certificates of Health Care Benefits, Benefit Booklets, Summary Plan Descriptions, and other coverage documents. Blue Cross and Blue Shield of Texas may use reasonable discretion interpreting and applying this policy to services being delivered in a particular case. BCBSTX has full and final discretionary authority for their interpretation and application to the extent provided under any applicable Plan documents.

Providers are responsible for submitting accurate documentation of services performed. Providers are expected to submit claims for services rendered using valid code combinations from Health Insurance Portability and Accountability Act approved code sets. Claims should be coded appropriately according to industry standard coding guidelines including, but not limited to: Uniform Billing Editor, American Medical Association, Current Procedural Terminology (CPT®) Assistant, Healthcare Common Procedure Coding System, ICD-10-CM and ICD-10-PCS, National Drug Codes, Diagnosis Related Group guidelines, Centers for Medicare & Medicaid Services National Correct Coding Initiative Policy Manual, CCI table edits and other CMS guidelines.

Claims are subject to the code edit protocols for services and procedures billed. Claim submissions are subject to claim review, including but not limited to, any terms of benefit coverage, provider contract language, medical policies, and reimbursement policies, as well as coding software logic. Upon request, the provider is urged to submit any additional documentation.

Description

The Plan has implemented certain lab management reimbursement criteria. Not all requirements apply to each product. Providers are urged to review Plan documents for eligible coverage for services rendered.

Reimbursement Information

NOTE: Except for where otherwise specified in the table below, quarterly measurement of designated serum biomarkers is permitted for follow-up, monitoring, and/or surveillance.

1) Measurement of the following serum biomarkers **may be reimbursable** for the following indications:

Serum Biomarker	Indication
Alkaline phosphatase (ALP)	<u>Bone neoplasms:</u> <ul style="list-style-type: none">• Workup for osteosarcoma
	<u>Melanoma (uveal):</u> <ul style="list-style-type: none">• Workup for distant metastatic disease
	<u>Systemic light chain amyloidosis:</u> <ul style="list-style-type: none">• Initial diagnostic workup to assess organ involvement for suspected SLCA
Alpha fetoprotein (AFP)	<u>Hepatocellular carcinoma:</u> <ul style="list-style-type: none">• Screening for at risk individuals due to Child-Turcotte-Pugh A or B cirrhosis or hepatitis B infection;• Workup for confirmed HCC;• Surveillance (every 3-6 months for 2 years, then every 6 months)
	<u>Intrahepatic cholangiocarcinoma:</u> <ul style="list-style-type: none">• Workup for isolated intrahepatic mass
	<u>Occult primary:</u> <ul style="list-style-type: none">• Additional workup for adenocarcinoma or carcinoma not otherwise specified with a clinical presentation in the liver or, mediastinum, or for individuals <65 years of age with a retroperitoneal mass;
	<u>Epithelial ovarian cancer/fallopian tube cancer/primary peritoneal cancer:</u> <ul style="list-style-type: none">• Initial workup for individuals with a suspicious mass found by palpation or imaging;• Monitoring prior to each cycle of chemotherapy;

	<ul style="list-style-type: none"> Monitoring/follow-up for patients not receiving treatment (every 2-4 months for 2 years; every 3-6 months for 3 years; annually after year 5) <p><u>Ovarian cancers (less common):</u></p> <ul style="list-style-type: none"> Initial workup for newly diagnosed ovarian cancer Monitoring/follow-up for recurrence <p><u>Less common ovarian cancers (additional surveillance recommendations):</u></p> <ul style="list-style-type: none"> <u>Malignant germ cell tumors:</u> <ul style="list-style-type: none"> Surveillance (every 2 months in year 1, every 3 months in year 2, every 6 months in year 3; year 4 and beyond, annual testing up to year 10) <u>Malignant sex cord stromal tumors:</u> <ul style="list-style-type: none"> Surveillance if clinically indicated. If done, frequency based on stage (e.g., 6-12 months if early-stage, low-risk disease; 4-6 months if high-risk disease) <p><u>Testicular cancer:</u></p> <ul style="list-style-type: none"> Initial diagnostic workup for a suspicious testicular mass; Post-diagnostic workup; Post-orchietomy Follow-up for non-seminoma (no more than every 2 months) Follow-up for pure seminoma (no more than every 3 months) <p><u>Thymomas and thymic carcinomas:</u></p> <ul style="list-style-type: none"> Initial evaluation for individuals with a mediastinal mass
<p>Beta-2 microglobulin (B2M)</p>	<p><u>B-cell lymphomas (classic follicular, diffuse large B-cell, HIV-related, lymphoblastic; mantle cell):</u></p> <ul style="list-style-type: none"> Additional workup for specified B-cell lymphomas <p><u>Castleman Disease:</u></p> <ul style="list-style-type: none"> Additional workup with established diagnosis <p><u>Chronic lymphocytic leukemia/small lymphocytic lymphoma:</u></p> <ul style="list-style-type: none"> Additional workup with established diagnosis; For prognostic and/or therapy determination <p><u>Multiple myeloma:</u></p> <ul style="list-style-type: none"> Initial diagnostic workup; <p><u>Systemic light chain amyloidosis:</u></p> <ul style="list-style-type: none"> Initial diagnostic workup to assess organ involvement for suspected SLCA <p><u>Waldenström macroglobulinemia / lymphoplasmacytic</u></p>

	<u>lymphoma:</u> <ul style="list-style-type: none"> • Additional workup with established diagnosis
BNP or NT-proBNP	<u>Multiple myeloma:</u> <ul style="list-style-type: none"> • Initial diagnostic workup
	<u>Systemic light chain amyloidosis:</u> <ul style="list-style-type: none"> • Initial diagnostic workup to assess organ involvement for suspected SLCA
Calcitonin (CALCA)	<u>Head and neck cancer – occult primary:</u> <ul style="list-style-type: none"> • Additional workup for squamous cell carcinoma, adenocarcinoma, and anaplastic undifferentiated epithelial tumors of the neck
	<u>Medullary carcinoma:</u> <ul style="list-style-type: none"> • Additional workup with established diagnosis; • Surveillance (every 6-12 months)
	<u>Multiple endocrine neoplasia, type 2:</u> <ul style="list-style-type: none"> • At diagnosis (clinical evaluation) to assess risk for medullary thyroid cancer
Cancer antigen 15-3 and 27.29 (CA 15-3 and 27.29)	<u>Breast cancer (invasive):</u> <ul style="list-style-type: none"> • Monitoring metastatic disease
	<u>Occult primary- suspected metastatic malignancy:</u> <ul style="list-style-type: none"> • Initial workup;
Cancer antigen 19-9 (CA 19-9)	<u>Ampullary adenocarcinoma:</u> <ul style="list-style-type: none"> • Additional workup with established diagnosis; • Surveillance for resected ampullary cancer (every 3-6 months for 2 years, every 6-12 months for up to 5 years as clinically indicated)
	<u>Appendiceal adenocarcinoma:</u> <ul style="list-style-type: none"> • Workup to establish baseline when imaging suggests appendiceal neoplasm or cancer or when there is a confirmed appendiceal neoplasm or cancer; • Workup for recurrence (3-6 months after CT imaging); • Surveillance (every 6-12 months for 2 years; annually up to 10 years).
	<u>Extrahepatic cholangiocarcinoma:</u> <ul style="list-style-type: none"> • Workup to establish baseline for individuals with abdominal pain, jaundice, abnormal liver function tests, or obstruction or abnormality on imaging
	<u>Gallbladder cancer:</u> <ul style="list-style-type: none"> • Workup to establish baseline for individuals with jaundice or a mass on imaging; • Post-operative workup of a suspicious mass; • Surveillance (as clinically indicated), post-resection

	<p><u>Intrahepatic cholangiocarcinoma:</u></p> <ul style="list-style-type: none"> • Workup to establish baseline for an isolated intrahepatic mass
	<p><u>Occult primary – suspected metastatic malignancy:</u></p> <ul style="list-style-type: none"> • Initial workup
	<p><u>Epithelial ovarian cancer/fallopian tube cancer/primary peritoneal cancer:</u></p> <ul style="list-style-type: none"> • Initial workup for individuals with a suspicious mass found by palpation or imaging; • Monitoring prior to each cycle of chemotherapy; • Monitoring/follow-up for patients not receiving treatment (every 2-4 months for 2 years; every 3-6 months for 3 years; annually after year 5)
	<p><u>Ovarian cancers (less common):</u></p> <ul style="list-style-type: none"> • Initial workup for newly diagnosed ovarian cancer; • Monitoring/follow-up for recurrence <p><u>Less common ovarian cancers (additional surveillance recommendations):</u></p> <ul style="list-style-type: none"> • <u>Malignant germ cell tumors:</u> <ul style="list-style-type: none"> ○ Surveillance (every 2 months in year 1; every 3 months in year 2; every 6 months in year 3; year 4 and beyond, annual testing up to year 10) • <u>Malignant sex cord stromal tumors:</u> <ul style="list-style-type: none"> ○ Surveillance if clinically indicated. If done, frequency based on stage (e.g., 6-12 months if early-stage, low-risk disease; 4-6 months if high-risk disease)
	<p><u>Pancreatic adenocarcinoma:</u></p> <ul style="list-style-type: none"> • Workup to establish baseline when there is either a clinical suspicion of pancreatic cancer with suspicious imaging or evidence of dilated pancreatic and/or bile duct (stricture) and when no metastatic disease is detected; • Post-operative, post-adjuvant treatment surveillance as appropriate
	<p><u>Small bowel adenocarcinoma:</u></p> <ul style="list-style-type: none"> • Additional workup to establish baseline with established diagnosis; • Post-treatment surveillance (every 3-6 months for 2 years, then every 6 months for a total of 5 years) • Additional workup for metastatic adenocarcinoma
<p>Cancer antigen 125 (CA-125)</p>	<p><u>Appendiceal adenocarcinoma:</u></p> <ul style="list-style-type: none"> • Workup to establish baseline when imaging suggests appendiceal neoplasm or cancer or when

	<p>there is a confirmed appendiceal neoplasm or cancer;</p> <ul style="list-style-type: none"> • Workup for recurrence (3-6 months after CT imaging); • Surveillance (every 6-12 months for 2 years; annually up to 10 years)
	<p><u>Endometrial carcinoma:</u></p> <ul style="list-style-type: none"> • Additional workup; • Surveillance (if initially elevated)
	<p><u>Occult primary:</u></p> <ul style="list-style-type: none"> • Initial workup for a suspected metastatic malignancy; • Additional workup for adenocarcinoma or carcinoma not otherwise specified (in those with a uterus and/or ovaries present) with a clinical presentation in the peritoneal/ascites or inguinal nodes
	<p><u>Epithelial ovarian cancer/fallopian tube cancer/primary peritoneal cancer:</u></p> <ul style="list-style-type: none"> • Initial workup for individuals with a suspicious mass found by palpation or imaging; • Monitoring prior to each cycle of chemotherapy; • Monitoring/follow-up for patients not receiving treatment (every 2-4 months for 2 years; every 3-6 months for 3 years; annually after year 5)
	<p><u>Ovarian cancers (less common):</u></p> <ul style="list-style-type: none"> • Initial workup for newly diagnosed ovarian cancer; • Monitoring/follow-up for recurrence <p><u>Less common ovarian cancers (additional surveillance recommendations):</u></p> <ul style="list-style-type: none"> • <u>Malignant germ cell tumors:</u> <ul style="list-style-type: none"> ○ surveillance (every 2 months in year 1; every 3 months in year 2; every 6 months in year 3; year 4 and beyond, annual testing up to year 10) • <u>Malignant sex cord stromal tumors:</u> <ul style="list-style-type: none"> ○ Surveillance if clinically indicated. If done, frequency based on stage (e.g., 6-12 months if early-stage, low-risk disease; 4-6 months if high-risk disease)

	<p><u>Uterine neoplasms</u></p> <ul style="list-style-type: none"> • Initial diagnostic workup for known or suspected malignancy; • Additional workup for serous, clear cell carcinoma, undifferentiated/dedifferentiated carcinoma, carcinosarcoma, suspected extrauterine disease; • Surveillance in endometrial carcinoma if initially elevated
<p>Carcinoembryonic antigen (CEA)</p>	<p><u>Appendiceal adenocarcinoma:</u></p> <ul style="list-style-type: none"> • Workup to establish baseline when imaging suggests appendiceal neoplasm or cancer or when there is a confirmed appendiceal neoplasm or cancer; • Workup for recurrence (3-6 months after CT imaging); • Surveillance (every 6-12 months for 2 years; annually up to 10 years)
	<p><u>Breast cancer (invasive):</u></p> <ul style="list-style-type: none"> • Monitoring metastatic disease
	<p><u>Colon cancer:</u></p> <ul style="list-style-type: none"> • Workup to establish baseline for a pedunculated or sessile polyp with invasive cancer, colon cancer appropriate for resection, and suspected or proven metastatic adenocarcinoma; • Surveillance (every 3-6 months for 2 years, then every 6 months for a total of 5 years)
	<p><u>Extrahepatic cholangiocarcinoma:</u></p> <ul style="list-style-type: none"> • Workup to establish baseline for individuals with abdominal pain, jaundice, abnormal liver function tests, or obstruction or abnormality on imaging
	<p><u>Gallbladder cancer:</u></p> <ul style="list-style-type: none"> • Workup to establish baseline for individuals with jaundice or a mass on imaging; • Post-operative workup of a suspicious mass; • Surveillance (as clinically indicated) post-resection
	<p><u>Intrahepatic cholangiocarcinoma:</u></p> <ul style="list-style-type: none"> • Workup to establish baseline for an isolated intrahepatic mass
	<p><u>Medullary carcinoma:</u></p> <ul style="list-style-type: none"> • Additional workup with established diagnosis; • Surveillance (every 6-12 months)

	<p><u>Multiple endocrine neoplasia, type 2:</u></p> <ul style="list-style-type: none"> • At diagnosis (clinical evaluation) to assess risk for medullary thyroid cancer
	<p><u>Epithelial ovarian cancer/fallopian tube cancer/primary peritoneal cancer:</u></p> <ul style="list-style-type: none"> • Initial workup; for individuals with a suspicious mass found by palpation or imaging; • Monitoring prior to each cycle of chemotherapy; • Monitoring/follow-up for patients not receiving treatment (every 2-4 months for 2 years; every 3-6 months for 3 years; annually after year 5)
	<p><u>Ovarian cancers (less common):</u></p> <ul style="list-style-type: none"> • Initial workup for newly diagnosed ovarian cancer; • Monitoring/follow-up for recurrence <p><u>Less common ovarian cancers (additional surveillance recommendations):</u></p> <ul style="list-style-type: none"> • <u>Malignant germ cell tumors:</u> <ul style="list-style-type: none"> ○ Surveillance (every 2 months in year 1; every 3 months in year 2, every 6 months in year 3; year 4 and beyond, annual testing up to year 10) • <u>Malignant sex cord stromal tumors:</u> <ul style="list-style-type: none"> ○ Surveillance if clinically indicated. If done, frequency based on stage (e.g., 6-12 months if early-stage, low-risk disease; 4-6 months if high-risk disease)
	<p><u>Rectal cancer:</u></p> <ul style="list-style-type: none"> • Workup to establish baseline for diagnosed rectal cancer or for pedunculated or sessile polyp with invasive cancer; • Surveillance (every 3-6 months for 2 years, then every 6 months for a total of 5 years)
	<p><u>Small bowel adenocarcinoma:</u></p> <ul style="list-style-type: none"> • Additional workup with established diagnosis; • Post-treatment surveillance (every 3-6 months for 2 years, then every 6 months for a total of 5 years) • Additional workup for metastatic adenocarcinoma
<p>Chorionic gonadotropin beta polypeptide (CGB3)</p>	<p><u>Gestational trophoblastic neoplasia:</u></p> <ul style="list-style-type: none"> • Initial workup for suspected GTN; • During and post treatment (no more than weekly); • Follow-up/surveillance (no more than monthly for 12 months)
	<p><u>Occult primary:</u></p> <ul style="list-style-type: none"> • Additional workup for adenocarcinoma or carcinoma not otherwise specified with a clinical

	<p>presentation in the mediastinum or for individuals < 65 years of age with a retroperitoneal mass</p> <p><u>Epithelial ovarian cancer/fallopian tube cancer/primary peritoneal cancer:</u></p> <ul style="list-style-type: none"> • Initial workup for individuals with a suspicious mass found on palpation or imaging; • Monitoring prior to each cycle of chemotherapy; • Monitoring/follow-up for patients not receiving treatment (every 2-4 months for 2 years; every 3-6 months for 3 years; annually after year 5) <p><u>Less common ovarian cancers:</u></p> <ul style="list-style-type: none"> • Initial workup for newly diagnosed ovarian cancer; • Monitoring/follow-up for recurrence • <u>Malignant germ cell tumors:</u> <ul style="list-style-type: none"> ○ Surveillance (every 2 months in year 1; every 3 months in year 2; every 6 months in year 3; year 4 and beyond, annual testing up to year 10) • <u>Malignant sex cord stromal tumors:</u> <ul style="list-style-type: none"> ○ Surveillance if clinically indicated. If done, frequency based on stage (e.g., 6-12 months if early-stage, low-risk disease; 4-6 months if high-risk disease) <p><u>Testicular cancer:</u></p> <ul style="list-style-type: none"> • Initial diagnostic workup for a suspicious testicular mass; • Post-diagnostic workup; • Post-orchietomy; • Follow-up for non-seminoma (no more than every 2 months) • Follow-up for pure seminoma (no more than every 3 months) <p><u>Thymomas and thymic carcinomas:</u></p> <ul style="list-style-type: none"> • Initial evaluation, for individuals with a mediastinal mass
<p>Human epididymis protein 4 (HE4)</p>	<p><u>Epithelial ovarian cancer/fallopian tube cancer/primary peritoneal cancer:</u></p> <ul style="list-style-type: none"> • Initial workup for individuals with a suspicious mass found on palpation or imaging; • Monitoring prior to each cycle of chemotherapy; • Monitoring/follow-up for patients not receiving treatment (every 2-4 months for 2 years; every 3-6 months for 3 years; annually after year 5) <p><u>Less common ovarian cancers:</u></p> <ul style="list-style-type: none"> • Initial workup for newly diagnosed ovarian cancer;

	<ul style="list-style-type: none"> Monitoring/follow-up for recurrence <p><u>Less common ovarian cancers (additional surveillance recommendations):</u></p> <ul style="list-style-type: none"> <u>Malignant germ cell tumors:</u> <ul style="list-style-type: none"> Surveillance (every 2 months in year 1; every 3 months in year 2; every 6 months in year 3; year 4 and beyond, annual testing up to year 10) <u>Malignant sex cord stromal tumors:</u> <ul style="list-style-type: none"> Surveillance if clinically indicated. If done, frequency based on stage (e.g., 6-12 months if early-stage, low-risk disease; 4-6 months if high-risk disease)
Inhibin (INHA)	<p><u>Epithelial ovarian cancer/fallopian tube cancer/primary peritoneal cancer:</u></p> <ul style="list-style-type: none"> Initial workup for individuals with a suspicious mass found on palpation or imaging; Monitoring prior to each cycle of chemotherapy; Monitoring/follow-up for patients not receiving treatment (every 2-4 months for 2 years; every 3-6 months for 3 years; annually after year 5) <p><u>Less common ovarian cancers:</u></p> <ul style="list-style-type: none"> Initial workup for newly diagnosed ovarian cancer; Monitoring/follow-up for recurrence <p><u>Less common ovarian cancers (additional surveillance recommendations):</u></p> <ul style="list-style-type: none"> <u>Malignant germ cell tumors:</u> <ul style="list-style-type: none"> Surveillance (every 2 months in year 1; every 3 months in year 2; every 6 months in year 3; year 4 and beyond, annual testing up to year 10) <u>Malignant sex cord stromal tumors:</u> <ul style="list-style-type: none"> Surveillance if clinically indicated. If done, frequency based on stage (e.g., 6-12 months if early-stage, low-risk disease; 4-6 months if high-risk disease)
Serum free light chains	<p><u>Castleman disease:</u></p> <ul style="list-style-type: none"> Additional workup for established diagnosis <p><u>Multiple myeloma:</u></p> <ul style="list-style-type: none"> Initial diagnostic workup; for multiple myeloma and polyneuropathy, organomegaly, endocrinopathy, monoclonal protein, skin changes (POEMS); Follow-up/surveillance for solitary plasmacytoma (every 3-6 months, asymptomatic smoldering myeloma (every 3-6 months), symptomatic multiple

	myeloma
	Systemic light chain amyloidosis: <ul style="list-style-type: none"> Initial diagnostic workup
Troponin T	Systemic light chain amyloidosis: <ul style="list-style-type: none"> Initial diagnostic workup to assess organ involvement for suspected SLCA
Tryptase	Systemic mastocytosis: <ul style="list-style-type: none"> Initial diagnosis

- 2) For all other cancer indications not discussed above, use of the above biomarkers (alone or in a panel of serum tumor markers) **are not reimbursable.**
- 3) All other serum tumor markers not addressed above (alone or in a panel of serum tumor markers) **are not reimbursable.**
- 4) For the screening and detection of cancer, analysis of proteomic patterns in serum **are not reimbursable.**

Procedure Codes

The following is not an all-encompassing code list. The inclusion of a code does not guarantee it is a covered service or eligible for reimbursement.

Code	Description
81500	ONCO (OVAR) TWO PROTEINS
81503	ONCO (OVAR) FIVE PROTEINS
81538	ONCOLOGY LUNG
82105	ALPHA-FETOPROTEIN SERUM
82107	ALPHA-FETOPROTEIN L3
82232	ASSAY OF BETA-2 PROTEIN
82308	ASSAY OF CALCITONIN
82378	CARCINOEMBRYONIC ANTIGEN
83520	IMMUNOASSAY QUANT NOS NONAB
83521	IG LIGHT CHAINS FREE EACH
83880	ASSAY OF NATRIURETIC PEPTIDE
83950	ONCOPROTEIN HER-2/NEU
83951	ONCOPROTEIN DCP
84075	ASSAY ALKALINE PHOSPHATASE
84484	ASSAY OF TROPONIN QUANT
84702	CHORIONIC GONADOTROPIN TEST
84704	HCG FREE BETACHAIN TEST
86300	IMMUNOASSAY TUMOR CA 15-3
86301	IMMUNOASSAY TUMOR CA 15-3
86304	IMMUNOASSAY TUMOR CA 125

86305	HUMAN EPIDIDYMIS PROTEIN 4
86316	IMMUNOASSAY TUMOR OTHER
86336	INHIBIN A
0003U	ONC OVAR 5 PRTN SER ALG SCOR
0092U	ONC LNG 3 PRTN BMRK PLSM ALG
0163U	ONC CLRCT SCR 3 PRTN ALG
0404U	ONC BRST SEMIQ MEAS THYM KN
0558U	ONC CLRCT ELISA BF7 AG SERUM
0559U	ONC BRS QUAN ELISA BF9AG SRM
0599U	ONC PNCRTC CA MULT IA SERUM
G0327	Colon ca scrn;bld-bsd biomrk

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Centers for Medicare & Medicaid Services. (2026). Healthcare Common Procedure Coding System (HCPCS) Level II.

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- NCCN. Biomarkers Compendium. <https://www.nccn.org/compendia-templates/compendia/biomarkers-compendium>
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- ASPIRA. OvaWatch. <https://aspirawh.com/ovawatch/>
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Approval Date	Description
05/15/2026	09/04/2026; Document updated with literature review. The following changes were made: Revisions made to indications in numerous biomarkers to reflect current recommendations in NCCN Biomarker's Compendium (e.g., for ALP testing for bone neoplasms, added "for osteosarcoma"). For BNP/NT-proBNP, added SLCA indications for testing. For calcitonin, CEA, and inhibin, removed "occult primary" recommendations, as the NCCN recommends these as an IHC marker for occult primary, not as a serum biomarker. For CA-125, removed "peritoneal mesothelioma" and "Lynch syndrome", as the NCCN no longer includes it as a recommended biomarker. Removed codes 81599, 83789, 84078, 84080, 84703, 84999. References revised.
07/25/2025	11/07/2025; Document updated with literature review. The following changes were made to Reimbursement Information: Changed "serum tumor markers" to "biomarkers" in initial note and in #1 to broaden definition as some serum-related markers are more accurately describes as biomarkers rather than serum tumor markers. Changed title in table to "Serum Biomarkers." Alkaline phosphatase (ALP): removed "during treatment" and "surveillance" for bone neoplasms. Added "Melanoma (uveal)" as an indication for workup for ALP. Alpha fetoprotein (AFP): Added: Carcinosarcoma (malignant mixed mullerian tumors): Monitoring; Follow-up; Clear cell carcinoma of the ovary: Monitoring; Follow - up; Mucinous neoplasms of the ovary: Monitoring; Follow-up; Low-grade serous carcinoma: Monitoring; Follow-up; Added header Ovarian Cancers; Removed "post-diagnostic" from Testicular cancer - nonseminoma to clarify workup can occur before or after diagnosis. Beta-2 microglobulin: For Beta-2 microglobulin, removed "Castleman disease" from B-cell lymphoma title and created a new row which identifies the indication for Castleman disease as "workup" with B2M measurement. For Chronic lymphocytic leukemia/small lymphocytic lymphoma, added For prognostic and/or therapy determination. Beta human chorionic gonadotropin (beta HCG) was renamed to Chorionic gonadotropin beta polypeptide (CGB3): Changed the words "testes presenting with" to "mediastinum or" under "Occult primary" designation. BNP or NT-proBNP: For BNP or NT-proBNP, removed "systemic light chain amyloidosis" and indication for "initial diagnostic workup" from BNP or NT-proBNP section (this was moved to a separate section with Troponin T.). Calcitonin (CALCA): added Adenocarcinoma and anaplastic/undifferentiated epithelial tumors: Workup; and Occult primary (unknown primary cancer): Workup. Cancer antigen 15-2 and 27.29: Added Occult primary: suspected

	<p>metastatic malignancy: Initial workup, assessing disease prognosis and monitoring/follow-up for response. Cancer antigen 19-9: Added Occult primary: workup to establish baseline; In "Ovarian cancers (less common)" merged ovarian cancer sections. Added "monitoring/follow up" as indications to "Carcinosarcoma, Clear cell carcinoma of the ovary, Grade 1 endometrial carcinoma, low-grade serous carcinoma," and "mucinous neoplasms of the ovary." In Small bowel adenocarcinoma, added "At metastasis or recurrence." Cancer antigen 125 (CA-125): For cancer antigen 125 (CA-125), added "initial evaluation/workup" to indications for Occult primary. Added "additional workup/surveillance" indications to uterine neoplasms. Carcinoembryonic antigen (CEA): For Carcinoembryonic antigen (CEA), added "Monitoring of adjuvant treatment" to Gallbladder cancer; Added Occult primary (unknown primary cancer) and indication for "workup for adenocarcinoma or carcinoma not otherwise specified." In "Ovarian cancers (less common)" merged ovarian cancer sections. Added "monitoring/follow up" as indications to "Carcinosarcoma, Clear cell carcinoma of the ovary, Grade 1 endometrial carcinoma, low-grade serous carcinoma," and "mucinous neoplasms of the ovary." Added "Post-adjuvant treatment" to Borderline epithelial tumors. Human epididymis protein 4 (HE4): For Human epididymis protein 4, added new section to the table. Added "Ovarian cancer/fallopian tube cancer/primary peritoneal cancer" with indications for "initial workup during primary chemotherapy; monitoring/follow-up for complete response (as clinically indicated)." Added "Ovarian cancers (less common) and indications for various cancers under this designation for "monitoring/follow-up." Added "Ovarian cancers" and additional indications for "borderline epithelial tumors." Added Inhibin (INHA) for Occult primary (unknown primary cancer); Ovarian cancer/fallopian tube cancer/primary peritoneal cancer; Ovarian cancers; Removed entire section on Lactate dehydrogenase (LDH) as LDH is a broad marker beyond serum tumor biomarker designation; Serum free light chains: Added "Castleman disease" with indication of workup; and added "follow-up" to Multiple myeloma. Added codes 0558U, 0559 effective 7/1/2025; added code 0599U effective 10/1/2025; removed code 83615. References revised.</p>
09/13/2024	01/01/2025: New policy