

# Local Coverage Determination (LCD): MolDX: Breast Cancer Index Genetic Assay (L36314)

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## Contractor Information

CONTRACTOR NAME	CONTRACT TYPE	CONTRACT NUMBER	JURISDICTION	STATE(S)
Noridian Healthcare Solutions, LLC	A and B MAC	01111 - MAC A	J - E	California - Entire State
Noridian Healthcare Solutions, LLC	A and B MAC	01112 - MAC B	J - E	California - Northern
Noridian Healthcare Solutions, LLC	A and B MAC	01182 - MAC B	J - E	California - Southern
Noridian Healthcare Solutions, LLC	A and B MAC	01211 - MAC A	J - E	American Samoa Guam Hawaii Northern Mariana Islands
Noridian Healthcare Solutions, LLC	A and B MAC	01212 - MAC B	J - E	American Samoa Guam Hawaii Northern Mariana Islands
Noridian Healthcare Solutions, LLC	A and B MAC	01311 - MAC A	J - E	Nevada
Noridian Healthcare Solutions, LLC	A and B MAC	01312 - MAC B	J - E	Nevada
Noridian Healthcare Solutions, LLC	A and B MAC	01911 - MAC A	J - E	American Samoa California - Entire State Guam Hawaii Nevada Northern Mariana Islands

## LCD Information

## Document Information

LCD ID

Original Effective Date

L36314

For services performed on or after 10/01/2015

**LCD Title**

MoIDX: Breast Cancer Index&#8480; Genetic Assay

**Revision Effective Date**

For services performed on or after 01/01/2019

**Proposed LCD in Comment Period**

N/A

**Revision Ending Date**

N/A

**Source Proposed LCD**

N/A

**Retirement Date**

N/A

**AMA CPT / ADA CDT / AHA NUBC Copyright Statement**

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**Notice Period Start Date**

N/A

**Notice Period End Date**

N/A

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**CMS National Coverage Policy**

Title XVIII of the Social Security Act, §1862(a)(1)(A) allows coverage and payment for only those services that are considered to be reasonable and necessary for the diagnosis or treatment of illness or injury or to improve the

functioning of a malformed body member

Title XVIII of the Social Security Act, §1862(a)(1)(D) items and services related to research and experimentation

Title XVIII of the Social Security Act, §1833(e), prohibits Medicare payment for any claim which lack the necessary information to process the claim.

42 CFR 410.32(a) Diagnostic x-ray tests, diagnostic laboratory tests, and other diagnostic tests: Conditions

42CFR411.15(k)(1) Particular services excluded from coverage

CMS On-Line Manual, Publication 100-08, Medicare Program Integrity Manual, Chapter 3, §3.4.1.3, diagnosis code requirements

## Coverage Guidance

### Coverage Indications, Limitations, and/or Medical Necessity

This policy limits coverage of the Breast Cancer Index (aka BCI) (bioTheranostics) to patients that meet the following criteria:

- Post-menopausal female with non-relapsed, ER+ breast cancer, **and**
- Was lymph node negative, **and**
- Is completing five (5) years of tamoxifen therapy, **and**
- Patient must be eligible for consideration of extended endocrine therapy based on published clinical trial data or practice guidelines, **and**
- Physician or patient is concerned about continuing anti-hormonal therapy because of documented meaningful toxicity or possible significant patient-specific side effects, **and**
- The test results will be discussed with the patient (including the limitations of the testing method, the risks and benefits of either continuing or stopping the therapy based on the test, and current cancer management guidelines).

Claims for BCI testing will be denied when testing does not meet all of the above criteria.

## Summary of Evidence

The body of evidence for the adjuvant endocrine treatment of ER+ breast cancer is continuing to evolve. Most recently, with the results of the ATLAS and aTTom trials, tamoxifen for an additional 5 years of adjuvant therapy has been added to the standard of care regimens as a NCCN recommendation to reduce late recurrences (defined as cancer recurring after 5 years of therapy).

The BCI assay (which was initially developed before ATLAS and aTTom data was available), uses PCR to interrogate selected proliferation-related and endocrine signaling-related genes, and may identify a subset of postmenopausal women who are at increased risk of late relapses for ER+ breast cancer and who may derive a greater benefit from extended hormone therapy. Current guidelines recommend adjuvant hormone therapy for postmenopausal patients with ER+ disease consisting of tamoxifen for 10 years (5 years initially and then strong consideration for an additional 5 years based on ATLAS and aTTom trials), aromatase Inhibitor (AI) for 5 years (ATAC), tamoxifen for 5 years followed by AI for 5 years (MA.17), or AI for 2-3 years followed by tamoxifen to complete 5 years. Extended (> 5 years) hormone therapy in each scenario is based on large randomized trials. Although there can be significant side

effects of these therapies in some women, generally the side effect profile is manageable when compared to the significant benefit of these interventions. The optimal sequence and duration of AI therapy, the benefit of tamoxifen after prolonged AI therapy, and result of tamoxifen use beyond 10 years remain unknown.

Although the BCI assay may identify a group of patients at highest risk of recurrence after 5 years post surgery it cannot be used independently without consideration of other clinical factors (such as age, tumor size and number of lymph nodes involved). A significant concern exists as to whether the assay can identify any group at such a low risk that it is safe to depart from a strong expert recommendation and safely stop therapy. Data from the prospective-retrospective review of the ATAC trial (trans-ATAC) shows a late recurrence risk in all patients, independent of risk category (BCI-H, BCI-I or BCI-L). Although the BCI-L group is lower than the others, it is still roughly 3.5% (but with 95% CI could be as high at 6.1%) and the annual risk of relapse steadily climbs in the years after stopping adjuvant endocrine therapy at 5 years. Given the substantial number of patients at risk in the BCI-Low group (over 60%), the total number of women at risk that may benefit from extended hormone therapy at the end of 5 years of tamoxifen therapy is not negligible.

Analysis of the prospective-retrospective review of MA.17 data, extended adjuvant therapy may be helpful in all groups studied. Although the H/I-low group did not meet statistical significance, the recurrence rate was decreased from 13% to 9% with the use of extended letrozole therapy with a relative risk reduction of 30%. In the H/I-High group the relative risk reduction was similar at 59%, which did meet statistical significance.

## **Analysis of Evidence (Rationale for Determination)**

### Level of Evidence

Quality: Moderate

Strength: Limited

Weight: Limited

The risk BCI-C model developed from the trans-ATAC data provides a continuous risk predicted by the test. The risk curve is flat in only the very lowest BCI values. Starting around BCI of 2 (in the middle of the BCI-Low category), there is a linear increase in recurrence risk. At the 95% confidence interval (CI), the risk in some individuals categorized in the BCI-low group could be as high as 20%. Due to the data complexity, there is a significant possibility that a physician might consider all BCI-L patients at negligible risk, and thus not consider extended hormone therapy and consequently lead women from the NCCN recommended interventions. Given the low toxicity and low cost of extended therapy, the false sense of security could deny many women from lifesaving therapy.

In women who have received an AI for 5 years, data is lacking on the utility of extended adjuvant hormonal therapy.

The data defined benefit of the BCI test appears to be when a woman is having significant side effects or has other concerns regarding adjuvant tamoxifen therapy and is opposed to taking more than 5 years of tamoxifen or starting on an AI (letrozole) after tamoxifen. If the toxicity or concern of extending hormone therapy is significant then it may be reasonable to use the BCI test to help make a risk/benefit decision with the patient on continued adjuvant endocrine therapy. In the majority of patients, the toxicity of extended hormone therapy is tolerable especially in

those who already have been on adjuvant hormonal therapy for 5 years. Even with the small benefit in all patients the MA.17, ATLAS and aTTom trials all demonstrated benefit from longer endocrine therapy. These trials established that the generally low toxicity of hormone therapy allows its long-term usage.

**BCI Use for Newly Diagnosed Breast Cancer Patients:**

It is possible that the BCI assay could identify patients who would most benefit from chemotherapy in the upfront setting, but this benefit has not been adequately studied. In one observational study the BCI risk recurrence level correlated with complete response to chemotherapy, but this trial does not provide sufficient evidence to warrant consideration for decision making in the newly diagnosed breast cancer patient. Further data will need to be published before the benefit of BCI can be confirmed or refuted in this setting.

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## Coding Information

**Bill Type Codes:**

Contractors may specify Bill Types to help providers identify those Bill Types typically used to report this service. Absence of a Bill Type does not guarantee that the policy does not apply to that Bill Type. Complete absence of all Bill Types indicates that coverage is not influenced by Bill Type and the policy should be assumed to apply equally to all claims.

CODE	DESCRIPTION
023x	Skilled Nursing - Outpatient

**Revenue Codes:**

Contractors may specify Revenue Codes to help providers identify those Revenue Codes typically used to report this service. In most instances Revenue Codes are purely advisory. Unless specified in the policy, services reported under other Revenue Codes are equally subject to this coverage determination. Complete absence of all Revenue Codes indicates that coverage is not influenced by Revenue Code and the policy should be assumed to apply equally to all Revenue Codes.

N/A

**CPT/HCPCS Codes**

**Group 1 Paragraph:**

N/A

**Group 1 Codes:**

CODE	DESCRIPTION
81518	Onc brst mrna 11 genes

**ICD-10 Codes that Support Medical Necessity**

**Group 1 Paragraph:**

N/A

**Group 1 Codes:**

ICD-10 CODE	DESCRIPTION
C50.011	Malignant neoplasm of nipple and areola, right female breast
C50.012	Malignant neoplasm of nipple and areola, left female breast
C50.019	Malignant neoplasm of nipple and areola, unspecified female breast
C50.111	Malignant neoplasm of central portion of right female breast
C50.112	Malignant neoplasm of central portion of left female breast
C50.119	Malignant neoplasm of central portion of unspecified female breast
C50.211	Malignant neoplasm of upper-inner quadrant of right female breast
C50.212	Malignant neoplasm of upper-inner quadrant of left female breast
C50.219	Malignant neoplasm of upper-inner quadrant of unspecified female breast
C50.311	Malignant neoplasm of lower-inner quadrant of right female breast
C50.312	Malignant neoplasm of lower-inner quadrant of left female breast
C50.319	Malignant neoplasm of lower-inner quadrant of unspecified female breast
C50.411	Malignant neoplasm of upper-outer quadrant of right female breast
C50.412	Malignant neoplasm of upper-outer quadrant of left female breast
C50.419	Malignant neoplasm of upper-outer quadrant of unspecified female breast
C50.511	Malignant neoplasm of lower-outer quadrant of right female breast
C50.512	Malignant neoplasm of lower-outer quadrant of left female breast
C50.519	Malignant neoplasm of lower-outer quadrant of unspecified female breast
C50.611	Malignant neoplasm of axillary tail of right female breast
C50.612	Malignant neoplasm of axillary tail of left female breast
C50.619	Malignant neoplasm of axillary tail of unspecified female breast
C50.811	Malignant neoplasm of overlapping sites of right female breast
C50.812	Malignant neoplasm of overlapping sites of left female breast
C50.819	Malignant neoplasm of overlapping sites of unspecified female breast
C50.911	Malignant neoplasm of unspecified site of right female breast
C50.912	Malignant neoplasm of unspecified site of left female breast
C50.919	Malignant neoplasm of unspecified site of unspecified female breast
Z17.0	Estrogen receptor positive status [ER+]

## ICD-10 Codes that DO NOT Support Medical Necessity

### Group 1 Paragraph:

BCI testing is non-covered for any indication other than those listed above.

**Group 1 Codes: N/A**

### Additional ICD-10 Information

N/A

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## General Information

### Associated Information

N/A

### Sources of Information

N/A

### Bibliography

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8. Strasser-Weippl K, et al. Extended adjuvant endocrine therapy in hormone-receptor positive breast cancer. *The Breast* 2013;22:S171-75.

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## Revision History Information

REVISION HISTORY DATE	REVISION HISTORY NUMBER	REVISION HISTORY EXPLANATION	REASON(S) FOR CHANGE
01/01/2019	R3	CPT code 81479 is deleted and replaced with 81518, per the 2019 Annual HCPCS Update.	<ul style="list-style-type: none"> <li>• Creation of Uniform LCDs With Other MAC Jurisdiction</li> <li>• Revisions Due To CPT/HCPCS Code Changes</li> </ul>
12/21/2017	R2	Added 21st Century Cures Act information	<ul style="list-style-type: none"> <li>• Creation of Uniform LCDs With Other MAC Jurisdiction</li> </ul>
11/17/2016	R1	This final LCD, effective 10/01/2015, combines JEA L36313 into the JEB LCD L36314 so that both JEA and JEB contract numbers will have the same final MCD LCD number.	<ul style="list-style-type: none"> <li>• Creation of Uniform LCDs Within a MAC Jurisdiction</li> </ul>

## Associated Documents

### Attachments

N/A

### Related Local Coverage Documents

Article(s)

A54214

- (MCD Archive Site)

### Related National Coverage Documents

N/A

### Public Version(s)

Updated on 12/19/2018 with effective dates 01/01/2019 - N/A

Updated on 12/12/2017 with effective dates 12/21/2017 - 12/31/2018

Some older versions have been archived. Please visit the MCD Archive Site to retrieve them.

## Keywords

- breast
- cancer
- genetic
- assay
- MoIDx
- Tamoxifen
- BCI



- bioTheranostics
- 81518