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Understanding a Breast Cancer

- Breast Cancer Stages
- Breast Cancer Survival Rates

Questions to Ask About Your Breast Cancer

You can take an active role in your breast cancer care by learning about your cancer and its treatment and by asking questions. Get a list of key questions here.

Questions to Ask Your Doctor About Breast Cancer

Connect with a breast cancer survivor

The American Cancer Society Reach To Recovery® program connects people facing breast cancer – from diagnosis through survivorship – with trained volunteers who are breast cancer survivors. Our volunteers provide one-on-one support through our website and mobile app to help those facing breast cancer cope with diagnosis, treatment, side effects, and more.

Breast Cancer Grade

Cancer cells are given a **grade** when they are removed from the breast and checked in the lab. Knowing a breast cancer's grade helps your cancer care team understand how

A **high grade number (grade 3)** means a faster-growing cancer that's more likely to spread.

An **intermediate grade number (grade 2)** means the cancer is growing faster than a grade 1 cancer but slower than a grade 3 cancer.

Grading invasive breast cancer cells

Three features of the invasive breast cancer cell are studied and each is given a score. The scores are then added to get a number between 3 and 9 that is used to get a grade of 1, 2, or 3, which is noted on your pathology report. Sometimes the terms *well differentiated*, *moderately differentiated*, and *poorly differentiated* are used to describe the grade instead of numbers:

- Grade 1 or well differentiated (score 3, 4, or 5). The cells are slower-growing, and look more like normal breast cells.
- Grade 2 or moderately differentiated (score 6, 7). The cells are growing at a speed of and look like cells somewhere between grades 1 and 3.
- **Grade 3 or poorly differentiated** (score 8, 9). The cancer cells look very different from normal cells and will probably grow and spread faster.

Our information about <u>pathology reports</u>¹ can help you understand details about your breast cancer.

Grading ductal carcinoma in situ (DCIS)

DCIS is also graded on how abnormal the cancer cells look and has a similar grading system to that used for invasive breast cancer (see above).

- **Grade 1 or low grade DCIS**. The cells are growing slower, and look more like normal breast cells. These cells tend to have estrogen and progesterone receptors (ER-positive and PR-positive).
- **Grade 2 or intermediate grade**. The cells are growing at a speed of and look like cells somewhere between grades 1 and 3.
- Grade 3 or high grade. The cancer cells look very different from normal cells and are growing faster. These cells tend not to have estrogen and progesterone receptors (ER-negative and PR-negative). High grade DCIS is often more likely to turn into invasive breast cancer.

Necrosis (areas of dead or dying cancer cells) is also noted. If there is necrosis, it means the tumor is growing quickly. The term *comedo necrosis* may be used if a breast duct is filled with dead and dying cells. Comedo necrosis is often linked to a high grade of DCIS and has a higher chance of developing into invasive breast cancer.

See <u>Understanding Your Pathology Report: Ductal Carcinoma In Situ</u>² for more on how DCIS is described.

Hyperlinks

- 1. <u>www.cancer.org/cancer/diagnosis-staging/tests/biopsy-and-cytology-tests/understanding-your-pathology-report.html</u>
- 2. www.cancer.org/cancer/diagnosis-staging/tests/biopsy-and-cytology-tests/understanding-your-pathology-report/breast-pathology/ductal-carcinoma-insitu.html

References

Bleiweiss IJ. Pathology of breast cancer. In Vora SR, ed. *UpToDate*. Waltham, Mass.: UpToDate, 2021. https://www.uptodate.com. Last updated June 12, 2020. Accessed August 31, 2021.

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Breast Cancer Ploidy and Cell Proliferation

Finding out more information about the DNA in breast cancer cells can help predict how fast those cancer cells are dividing and growing. If you want to learn more about DNA and breast cancer, you should first understand two important terms: **ploidy** and **cell proliferation**.

- What is ploidy and what does it mean?
- What is cell proliferation?

What is ploidy and what does it mean?

The **ploidy** of cancer cells refers to the amount of DNA they contain.

- If there's a normal amount of DNA in the cells, they are said to be **diploid**. These cancers tend to grow and spread more slowly.
- If the amount of DNA is abnormal, then the cells are called **aneuploid**. These cancers tend to be more aggressive. They also tend to grow and spread faster.

Tests of ploidy may help figure out long-term outcomes, but they rarely change treatment and are considered optional. They are not usually recommended as part of a routine breast cancer work-up.

What is cell proliferation?

Cell proliferation is how quickly a cancer cell copies its DNA and divides into 2 cells. If the cancer cells are dividing more rapidly, it means the cancer is faster growing or more aggressive.

The rate of cancer cell proliferation can be estimated by doing a **Ki-67 test**. In some cases, Ki-67 testing to measure cell proliferation may be used to help plan treatment or estimate treatment outcomes. But test results can vary depending on things like the lab doing the testing, the testing method, and what part of the tumor is tested.

Another way to determine cell division is the **S-phase fraction**, which is the percentage of cells in a sample that are copying their DNA as it gets ready to divide into 2 new cells.

If the S-phase fraction or Ki-67 labeling index is high, it means that the cancer cells are dividing more rapidly.

References

Duffy MJ, Harbeck N, Nap M, Molina R, Nicolini A, Senkus E, Cardoso F. Clinical use of biomarkers in breast cancer: Updated guidelines from the European Group on Tumor Markers (EGTM). *Eur J Cancer*. 2017;75:284-298.

Harris L, Fritsche H, Mennel R, et al. American Society of Clinical Oncology 2007 update of recommendations for the use of tumor markers in breast cancer. *J Clin Oncol.* 2007;25(33):5287-5312. doi:10.1200/JCO.2007.14.2364.

Jagsi R, King TA, Lehman C, Morrow M, Harris JR, Burstein HJ. Chapter 79: Malignant Tumors of the Breast. In: DeVita VT, Lawrence TS, Lawrence TS, Rosenberg SA, eds. *DeVita, Hellman, and Rosenberg's Cancer: Principles and Practice of Oncology*. 11th ed. Philadelphia, Pa: Lippincott Williams & Wilkins; 2019.

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Breast Cancer Hormone Receptor Status

Breast cancer cells taken out during a biopsy or surgery are tested to see if they have certain proteins that are **estrogen or progesterone receptors**. If your breast cancer cells have these receptors (proteins), this means that when the hormones estrogen and progesterone attach to the receptors, they stimulate the cancer to grow.

Cancers are called **hormone receptor-positive** or **hormone receptor-negative** based on whether or not they have these receptors (proteins).

Knowing the hormone receptor status of a cancer is important, because it helps determine the treatment options. Ask your cancer care team about your hormone receptor status and what it means for you.

- What are estrogen and progesterone receptors?
- Why is knowing hormone receptor status important?
- How are breast tumors tested for hormone receptors?
- What do the hormone receptor test results mean?

What are estrogen and progesterone receptors?

Receptors are proteins in or on cells that can attach to certain substances in the blood. Normal breast cells and some breast cancer cells have receptors that attach to the hormones estrogen and progesterone, and need these hormones for the cells to grow.

Breast cancer cells may have one, both, or none of these receptors.

- ER-positive: Breast cancers that have estrogen receptors are called ER-positive (or ER+) cancers.
- **PR-positive**: Breast cancers with progesterone receptors are called PR-positive (or **PR+**) cancers.
- Hormone receptor-positive: If the cancer cell has one or both of the receptors above, the term hormone-receptive positive (also called hormone-positive or HR+) breast cancer may be used.
- Hormone receptor-negative: If the cancer cell does not have the estrogen or the progesterone receptor, it's called hormone-receptor negative (also called hormonenegative or HR-).

Keeping the hormones estrogen and progesterone from attaching to the receptors can help keep the cancer from growing and spreading. There are <u>drugs that can be used to do this</u>¹.

Why is knowing hormone receptor status important?

Knowing the hormone receptor status of your cancer helps doctors decide how to treat it. If your cancer has one or both of these hormone receptors, hormone therapy drugs can be used to either lower estrogen levels or stop estrogen from acting on breast cancer cells. This kind of treatment is helpful for hormone receptor-positive breast cancers, but it doesn't work on tumors that are hormone receptor-negative (both ER-and PR-negative).

All invasive breast cancers should be tested for both of these hormone receptors either on the biopsy sample or when the tumor is removed with surgery. About 3

of 4 breast cancers have at least one of these receptors. This percentage is higher in older women than in younger women. DCIS should also be checked for hormone receptors.

How are breast tumors tested for hormone receptors?

A test called an **immunohistochemistry (IHC) test** is used most often to find out if cancer cells have estrogen and progesterone receptors. The test results will help guide you and your cancer care team in making the best treatment decisions.

What do the hormone receptor test results mean?

Test results will give you your hormone receptor status. It will say a tumor is hormone receptor-positive if at least 1% of the cells tested have estrogen and/or progesterone receptors. Otherwise, the test will say the tumor is hormone receptor-negative.

Hormone receptor-positive

receptor-positive breast cancer. In Vora SR, ed. *UpToDate*. Waltham, Mass.: UpToDate, 2021. https://www.uptodate.com. Last updated August 13, 2021. Accessed August 31, 2021.

Rimawi MF and Osborne CK. Chapter 43: Adjuvant Systemic Therapy: Endocrine Therapy. In: Harris JR, Lippman ME, Morrow M, Osborne CK, eds. *Diseases of the Breast*. 5th ed. Philadelphia: Wolters Kluwer Health; 2014.

Stearns V and Davidson NE. Chapter 45: Adjuvant Chemo Endocrine Therapy. In: Harris JR, Lippman ME, Morrow M, Osborne CK, eds. *Diseases of the Breast*. 5th ed. Philadelphia: Wolters Kluwer Health; 2014.

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Breast Cancer HER2 Status

About 15% to 20% of breast tumors have higher levels of a protein known as HER2. These cancers are called **HER2-positive breast cancers**. Ask your cancer care team about your HER2 status and what it means for you.

- What is HER2 and what does it mean?
- How are breast tumors tested for HER2?
- What do the test results mean?

What is HER2 and what does it mean?

HER2 is a protein that helps breast cancer cells grow quickly. Breast cancer cells with higher than normal levels of HER2 are called **HER2-positive**. These cancers tend to grow and spread faster than breast cancers that are HER2-negative, but are much more likely to respond to treatment with <u>drugs that target the HER2 protein</u>¹.

All invasive breast cancers should be tested for HER2 either on the biopsy sample or when the tumor is removed with surgery.

How are breast tumors tested for HER2?

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Hyperlinks

- 1. <u>www.cancer.org/cancer/types/breast-cancer/treatment/targeted-therapy-for-breast-cancer.html</u>
- 2. www.cancer.org/cancer/diagnosis-staging/tests/biopsy-and-cytology-tests.html
- 3. www.cancer.org/cancer/diagnosis-staging/tests/biopsy-and-cytology-tests/understanding-your-pathology-report/breast-pathology/breast-cancer-pathology.html
- 4. <u>www.cancer.org/cancer/types/breast-cancer/treatment/targeted-therapy-for-breast-cancer.html</u>

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Breast Cancer Gene Expression Tests

Gene expression tests are a form of <u>personalized medicine</u>¹. Personalized medicine is a way to learn more about your cancer and tailor your treatment.

These tests are done on breast cancer cells after surgery or biopsy to look at the patterns of a number of different genes. This process or test is sometimes called **gene expression profiling.**

- What do the test results mean?
- Testing options

What do the test results mean?

The patterns found can help predict if certain early-stage breast cancers are likely to come back after initial treatment.

Some gene expression testing/profiling can help predict which women will most likely benefit from chemotherapy² after breast surgery (adjuvant chemotherapy.) Hormone therapy³ is a standard treatment for hormone receptor-positive breast cancers, but it's not always clear when to use chemotherapy. These tests can help guide that decision. Still, these tests cannot tell any one woman for certain if her cancer will come back with or without chemotherapy.

These tests continue to be studied in large <u>clinical trials</u>⁴ to better understand how and when to best use them. In the meantime, ask your doctor if these tests might be useful for you.

Testing options

The Oncotype DX, MammaPrint, and Prosigna are examples of tests that look at different sets of breast cancer genes to see if chemotherapy is needed to help reduce the risk of cancer coming back (recurrence). More tests are in development. The type of test that's used will depend on your situation. Keep in mind that these tests are used for early-stage cancers, and testing isn't needed in all cases. For example, if breast cancer is advanced, it might be clear that chemotherapy is needed, even without gene

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option might be <u>ovarian suppression along with tamoxifen or an aromatase</u> inhibitor⁶.

• A high score (26-100) means a higher risk of recurrence. Women in this group are more likely to benefit from the addition of chemotherapy to hormone therapy to help lower the chance of the cancer coming back.

MammaPrint

The MammaPrint test can be used to help determine how likely breast cancers are to recur in a distant part of the body after treatment. It can be used for any type of invasive breast cancer that's 5cm (about 2 inches) or smaller and has spread to no more than 3 lymph nodes. This test can be done regardless of a woman's age or the cancer's hormone or HER2 status.

Hyperlinks

- 1. www.cancer.org/cancer/managing-cancer/treatment-types/precision-medicine.html
- 2. <u>www.cancer.org/cancer/types/breast-cancer/treatment/chemotherapy-for-breast-cancer.html</u>
- 3. <u>www.cancer.org/cancer/types/breast-cancer/treatment/hormone-therapy-for-breast-cancer.html</u>
- 4. <u>www.cancer.org/cancer/managing-cancer/making-treatment-decisions/clinical-trials.html</u>
- 5. <u>www.cancer.org/cancer/types/breast-cancer/about/types-of-breast-cancer/dcis.html</u>
- 6. <u>www.cancer.org/cancer/types/breast-cancer/treatment/hormone-therapy-for-breast-cancer.html</u>

References

Cardoso F, van't Veer LJ, Bogaerts J, Slaets L, Viale G, Delaloge S et al. 70-Gene Signature as an Aid to Treatment Decisions in Early-Stage Breast Cancer. *N Engl J Med*. 2016;375(8):717-29.

Foukakis T, and Bergh J, and Hurvitz SA. Deciding when to use adjuvant chemotherapy for hormone receptor-positive, HER2-negative breast cancer. In Vora SR, ed. *UpToDate*. Waltham, Mass.: UpToDate, 2021. https://www.uptodate.com. Last updated August 17, 2021. Accessed September 14, 2021.

Gnant M, Filipits M, Dubsky P, et al. Predicting risk for late metastasis: The PAM50 risk of recurrence (ROR) score after 5 years of endocrine therapy in postmenopausal women with HR+ early breast cancer: A study on 1,478 patients for the ABCSG-8 trial. *Ann Oncol.* 2013; 24(Suppl 3): iii29-iii37.

Harris LN, Ismaila N, McShane LM, et al. Use of Biomarkers to Guide Decisions on Adjuvant Systemic Therapy for Women With Early-Stage Invasive Breast Cancer: American Society of Clinical Oncology Clinical Practice Guideline. *J Clin Oncol*. 2016;34(10):1134–1150.

Knauer M, Mook S, Rutgers EJ, Bender RA, Hauptmann M, van de Vijver MJ et al. The

predictive value of the 70-gene signature for adjuvant chemotherapy in early breast cancer. *Breast Cancer Res Treat.* 2010 Apr;120(3):655-61.

Krop I, Ismaila N, Andre F, et al. Use of Biomarkers to Guide Decisions on Adjuvant Systemic Therapy for Women With Early-Stage Invasive Breast Cancer: American Society of Clinical Oncology Clinical Practice Guideline Focused Update. *J Clin Oncol.* 2017;35(24):2838–2847.

National Comprehensive Cancer Network (NCCN). Practice Guidelines in Oncology: Breast Cancer. Version 8.2021. Accessed at https://www.nccn.org/professionals/physician_gls/pdf/breast.pdf on Sept. 14, 2021.

Paik, S. Development and Clinical Utility of a 21-Gene Recurrence Score Prognostic Assay in Patients with Early Breast Cancer Treated with Tamoxifen. *The Oncologist*. 2007;12(6): 631-635.

Sparano JA, Gray RJ, Makower DF, Pritchard KI, Albain DF, Hayes DF, et al. Adjuvant chemotherapy guided by a 21-gene expression assay in breast cancer. *N Engl J Med.* 2018;379(2):111-121.

Sparano JA, Gray RJ, Ravdin PM, Makower DF, Pritchard KI, Albain KS et al. Clinical and Genomic Risk to Guide the Use of Adjuvant Therapy for Breast Cancer. *N Engl J Med*. 2019;380(25):2395-2405.

Wallden B, Storhoff J, Nielsen T, et al. Development and verification of the PAM50-based Prosigna breast cancer gene signature assay. *BMC Med Genomics*. 2015;8:54.

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Other Breast Cancer Gene, Protein, and Blood Tests

When breast cancer is diagnosed, samples that have been collected during <u>biopsies</u>¹, bloodwork, or other tests are sent to a pathology lab. A pathologist (a doctor who uses lab tests to diagnose diseases such as cancer) will look at the samples and may do

other special tests to help better classify the cancer.

These tests can also help choose certain drugs that might work better for your cancer. This is sometimes called <u>precision or personalized medicine</u>² because it is precise (or specific) for the features of your cancer.

The results of these tests are described in a pathology report, which is usually available within a week or two. If you have any questions about your pathology results or any diagnostic tests, talk to your doctor. If needed, you can get a second opinion of your pathology report by having your tissue samples sent to a pathologist at another lab.

- Tests for certain proteins on tumor cells
- Molecular tests for gene changes
- Blood tests

Tests for certain proteins on tumor cells

Lab tests might be done to look for certain proteins on the cancer cells.

Hormone receptor proteins: All breast cancers are tested for hormone receptors (proteins). Specifically, the cancer is tested for estrogen receptor (ER) and progesterone receptor (PR). Read more at Breast Cancer Hormone Receptor Status.

HER2 protein: All invasive breast cancers are tested for the HER2 protein to see if too much is being made. If it is not clear how much HER2 protein is present, molecular testing might be done to see how many copies of the *HER2* gene the cancer cells have. For more information about the HER2 gene and protein see Breast Cancer HER2 Status.

PD-L1 protein: People with advanced or metastatic triple-negative breast cancer might have their cancer tissue tested for the **PD-L1** protein, which can show if the cancer is more likely to respond to treatment with certain <u>immunotherapy</u>³ drugs along with chemotherapy.

Molecular tests for gene changes

In some cases, doctors may test for specific gene changes in the breast cancer cells that could mean certain <u>targeted drugs</u>⁴ or immunotherapy drugs might help treat the cancer.

These molecular tests (also known as **genomic tests** or **biomarker tests**) can be done on tissue taken during a biopsy or surgery for breast cancer. If the biopsy sample is too small and all the molecular tests cannot be done, the testing may also be done on blood that is taken from a vein just like a regular blood draw. This blood contains the DNA from dead tumor cells (known as **circulating tumor DNA**, or **ctDNA**). Obtaining the tumor DNA through a blood draw is sometimes called a "liquid biopsy" and can have advantages over a standard needle biopsy, which can carry risks.

Some changes that might be tested for include:

- BRCA1 and BRCA2 gene mutations: For women with an advanced HER2-negative breast cancer, your doctor might test you (not your cancer cells) for a hereditary BRCA1 or BRCA2 mutation⁵ (gene change). If you have one of these gene changes, treatment with the targeted drugs, olaparib (Lynparza) or talazoparib (Talzenna) might be options.
- PIK3CA, AKT1, and PTEN gene mutations: These 3 genes code for proteins in
 the same signaling pathway inside cells that can help them grow. Cancer cells
 sometimes have changes in one of these genes. If you have advanced breast
 cancer that is hormone receptor-positive and HER2-negative, and if the cancer
 cells show changes in one of these genes, a targeted drug such as capivasertib
 (Truqap) or alpelisib (Piqray) might be an option, along with the hormone drug⁶
 fulvestrant.
- *ESR1 gene mutations:* The *ESR1* gene contains the cell's instructions for the estrogen receptor (ER) protein. Mutations in this gene can make breast cancers less likely to be helped by some forms of hormone therapy. But for advanced breast cancer, the hormone drug elacestrant (Orserdu) may be helpful if the cancer cells have an *ESR1* mutation. This gene change can be tested for in a sample of your blood.
- MSI and MMR testing: Breast cancer cells might be tested to see if they show high levels of gene changes called microsatellite instability (MSI). Testing might also be done to see if the cancer cells have changes in any of the mismatch repair (MMR) genes (MLH1, MSH2, MSH6, and PMS2). Breast cancer cells that have a high level of microsatellite instability (MSI-H) or a defect in a mismatch repair gene (dMMR) might be treated with the immunotherapy drugs, pembrolizumab (Keytruda) or dostarlizumab (Jemperli).
- Tumor mutational burden (TMB): TMB is a measure of the number of gene mutations (changes) inside the cancer cells. Breast cancer cells that have many gene mutations (a high TMB) might be more likely to be recognized as abnormal and attacked by the body's immune system. If your breast cancer tissue is tested

and found to have a **high TMB (TMB-H)**, treatment with pembrolizumab (Keytruda) might be an option.

NTRK gene changes: Some breast cancer cells might have changes in one of the NTRK genes. These gene changes can sometimes lead to cancer growth.
 Larotrectinib (Vitrakvi) and entrectinib (Rozlytrek) are drugs that target the proteins made by the abnormal NTRK genes and might be options for people with advanced breast cancer.

Blood tests

Blood tests are not used to diagnose breast cancer, but they can help to get a sense of a person's overall health. For example, they can be used to help determine if a person is healthy enough to have surgery or certain types of chemotherapy.

A **complete blood count (CBC)** looks at whether your blood has normal numbers of different types of blood cells. For example, it can show if you are anemic (have a low number of red blood cells), if you could have trouble with bleeding (due to a low number of blood platelets), or if you are at increased risk for infections (because of a low number of white blood cells). This test could be repeated regularly during treatment, as many cancer drugs can affect blood-forming cells of the bone marrow.

Blood chemistry tests can help find if some of your organs, such as the liver or kidneys are not working as well. For example, if cancer has spread to the bones, it might cause higher than normal levels of calcium and alkaline phosphatase. If breast cancer spreads to the liver, it can sometimes cause high levels of liver function tests, such as aspartate aminotransferase (AST) or alanine aminotransferase (ALT). Breast cancer does not spread to the kidneys, but if your bloodwork shows your kidneys are not working well, certain chemo drugs, like cisplatin, might not be used.

Breast cancer cells sometimes make substr cebreast cancer.

Like CT scans, MRI scans³ show detailed images of soft tissues in the body. But MRI scans use radio waves and strong magnets instead of x-rays. This test can be used to look at the breasts or other parts of the body, such as the brain or spinal cord to look for possible cancer spread.

Ultrasound

<u>Ultrasound</u>⁴ (ultrasonography) uses sound waves to create an image on a video screen. A small microphone-like instrument called a transducer that gives off sound waves is moved over the skin surface and picks up the echoes as they bounce off tissues. A computer turns these echoes into an image on the screen. An ultrasound can be done over a breast or in the underarm area, or even the liver.

Positron emission tomography (PET) scan

For a <u>PET scan</u>⁵, a slightly radioactive form of sugar (known as FDG) is injected into the blood and collects mainly in cancer cells.

PET/CT scan: Often a PET scan is combined with a CT scan using a special machine that can do both at the same time. This lets the doctor compare areas of higher radioactivity on the PET scan with a more detailed picture on the CT scan.

Bone scan

A <u>bone scan</u>⁶ can help show if the cancer has spread to your bones. A small amount of low-level radioactive material is injected into the blood and collects mainly in abnormal areas of bone. It can show all of the bones of your body at the same time and can find small areas of cancer spread not seen on plain x-ray.

Hyperlinks

- 1. <u>www.cancer.org/cancer/diagnosis-staging/tests/imaging-tests/imaging-radiology-tests-for-cancer.html</u>
- 2. <u>www.cancer.org/cancer/diagnosis-staging/tests/imaging-tests/ct-scan-for-cancer.html</u>
- 3. www.cancer.org/cancer/diagnosis-staging/tests/imaging-tests/mri-for-cancer.html

- 4. www.cancer.org/cancer/diagnosis-staging/tests/imaging-tests/ultrasound-for-cancer.html
- 5. <u>www.cancer.org/cancer/diagnosis-staging/tests/imaging-tests/nuclear-medicine-scans-for-cancer.html</u>
- 6. <u>www.cancer.org/cancer/diagnosis-staging/tests/imaging-tests/nuclear-medicine-scans-for-cancer.html</u>

References

Joe BN. Clinical features, diagnosis, and staging of newly diagnosed breast cancer. In Vora SR, ed. *UpToDate*. Waltham, Mass.: UpToDate, 2021. https://www.uptodate.com. Last updated May 12, 2021. Accessed August 31, 2021.

National Comprehensive Cancer Network. NCCN Clinical Practice Guidelines in Oncology. Breast Cancer. Version 7.2021. Accessed at www.nccn.org/professionals/physician_gls/pdf/breast.pdf on August 31, 2021.

Niravath P, Osborne CK. Chapter 31: Evaluation of Patients for Metastasis Prior to Primary Therapy. In: Harris JR, Lippman ME, Morrow M, Osborne CK, eds. *Diseases of the Breast*. 5th ed. Philadelphia, Pa: Lippincott Williams & Wilkins; 2014.

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Breast Cancer Stages

After someone is diagnosed with breast cancer, doctors will try to figure out if it has spread, and if so, how far. This process is called **staging**. The stage of a cancer describes how much cancer is in the body. It helps determine how serious the cancer is and how best to <u>treat</u>¹ it. Doctors also use a cancer's stage when talking about <u>survival</u> statistics.

- What are the breast cancer stages?
- How is the stage determined?

Details of the TNM staging system

progesterone receptor?

- HER2 status: Does the cancer make too much of a protein called HER2?
- Grade of the cancer (G): How much do the cancer cells look like normal cells?

In addition, Oncotype Dx® Recurrence Score results may also be considered in the stage in certain situations.

Once all of these factors have been determined, this information is combined in a process called **stage grouping** to assign an overall stage. For more information see <u>Cancer Staging</u>².

Details about the first three factors (the TNM categories) are below. However, the addition of information about ER, PR, and HER2 status along with grade has made stage grouping for breast cancer more complex than for other cancers. Because of this, it is best to ask your doctor about your specific stage and what it means.

Details of the TNM staging system

Numbers or letters after T, N, and M provide more details about each of these factors. Higher numbers mean the cancer is more advanced. The categories below use the pathologic (surgical) definitions.

T categories for breast cancer

T followed by a number from 0 to 4 describes the main (primary) tumor's size and if it has spread to the skin or to the chest wall under the breast. Higher T numbers mean a larger tumor and/or wider spread to tissues near the breast.

TX: Primary tumor cannot be assessed.

T0: No evidence of primary tumor.

Tis: Carcinoma in situ (DCIS, or Paget disease of the breast with no associated tumor mass)

T1 (includes T1a, T1b, and T1c): Tumor is 2 cm (3/4 of an inch) or less across.

T2: Tumor is more than 2 cm but not more than 5 cm (2 inches) across.

T3: Tumor is more than 5 cm across.

T4 (includes T4a, T4b, T4c, and T4d) /GS1198 g

found in internal mammary lymph nodes (those near the breast bone) on sentinel lymph node biopsy.

N1mi:

OR

Cancer has spread to 4 or more axillary lymph nodes (with at least one area of cancer spread greater than 2 mm), and to the internal mammary lymph nodes on sentinel lymph node biopsy.

N3c: Cancer has spread to the lymph nodes above the collarbone (supraclavicular nodes) on the same side of the cancer with at least one area of cancer spread greater than 2 mm.

M categories for breast cancer

M followed by a 0 or 1 indicates whether the cancer has spread to distant organs -- for example, the lungs, liver, or bones.

M0: No distant spread is found on x-rays (or other imaging tests) or by physical exam.

cM0(i+): Small numbers of cancer cells are found in blood or bone marrow (found only by special tests), or tiny areas of cancer spread (no larger than 0.2 mm) are found in lymph nodes away from the underarm, collarbone, or internal mammary areas.

M1: Cancer has spread to distant organs (most often to the bones, lungs, brain, or liver) as seen on imaging tests or by physical exam, and/or a biopsy of one of these areas proves cancer has spread and is larger than 0.2mm.

Examples using the full staging system

Because there are so many factors that go into stage grouping for breast cancer, it's not possible to describe here every combination that might be included in each stage. The many different possible combinations mean that two women who have the same stage of breast cancer might have different factors that make up their stage.

Here are 3 examples of how all of the factors listed above are used to determine the pathologic (surgical) breast cancer stage:

Example #1

If the cancer size is between 2 and 5 cm (T2) but it has not spread to the nearby lymph nodes (N0) or to distant organs (M0) **AND** is:

Grade 3

- HER2 negative
- ER positive
- PR positive

The cancer stage is IB.

Example #2

If the cancer is larger than 5 cm (T3) and has spread to 4 to 9 lymph nodes under the arm or to any internal mammary lymph nodes (N2) but not to distant organs (M0) **AND** is:

- Grade 2
- HER2 positive
- ER positive
- PR positive

The cancer stage is IB.

Example #3

If the cancer is larger than 5 cm (T3) and has spread to 4 to 9 lymph nodes under the arm or to any internal mammary lymph nodes (N2) but not to distant organs (M0) **AND** is:

- Grade 2
- HER2 negative
- ER negative
- PR negative

The cancer stage is IIIB.

These are only 3 examples out of many possible combinations of factors. To understand what your breast cancer stage is, and what it means, talk to your doctor.

Hyperlinks

- 1. www.cancer.org/cancer/types/breast-cancer/treatment.html
- 2. www.cancer.org/cancer/diagnosis-staging/staging.html

References

American Joint Committee on Cancer. Breast. In: AJCC Cancer Staging Manual. 8th ed. New York, NY: Springer; 2017:589.

National Comprehensive Cancer Network (NCCN). Practice Guidelines in Oncology: Breast Cancer. Version 8.2021. Accessed at https://www.nccn.org/professionals/physician_gls/pdf/breast.pdf on September 15, 2021.

Paik, S. Development and Clinical Utility of a 21-Gene Recurrence Score Prognostic Assay in Patients with Early Breast Cancer Treated with Tamoxifen. *The Oncologist*. 2007;12(6): 631-635.

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Survival Rates for Breast Cancer

Survival rates can give you an idea of what percentage of people with the same type and stage of cancer are still alive a certain amount of time (usually 5 years) after they were diagnosed. They can't tell you how long you will live, but they may help give you a better understanding of how likely it is that your treatment will be successful.

Keep in mind that survival rates are estimates and are often based on previous outcomes of large numbers of people who had a specific cancer, but they can't predict what will happen in any particular person's case. These statistics can be confusing and may lead you to have more questions. Talk with your doctor, who is familiar with your situation, about how these numbers may apply to you.

What is a 5-year relative survival rate?

- Where do these numbers come from?
- 5-year relative survival rates for breast cancer
- Understanding the numbers

What is a 5-year relative survival rate?

Distant	31%
All SEER stages combined	91%

^{*}Localized stage only includes invasive cancer. It does not include ductal carcinoma in situ (DCIS).

Understanding the numbers

- Women now being diagnosed with breast cancer may have a better outlook than these numbers show. Treatments improve over time, and these numbers are based on women who were diagnosed and treated at least five years earlier.
- These numbers apply only to the stage of the cancer when it is first diagnosed. They do not apply later on if the cancer grows, spreads, or comes back after treatment.
- These numbers don't take everything into account. Survival rates are grouped based on how far the cancer has spread, but your age, overall health, how well the cancer responds to treatment, tumor grade, the presence of hormone receptors on the cancer cells, HER2 status, and other factors can also affect your outlook.
- Survival rates for women with triple-negative breast cancer are different than those above. See Triple-negative Breast Cancer¹.
- Survival rates for women with inflammatory breast cancer are different than those above. See<u>Inflammatory Breast Cancer</u>².

Hyperlinks

- 1. <u>www.cancer.org/cancer/types/breast-cancer/about/types-of-breast-cancer/triple-negative.html</u>
- 2. <u>www.cancer.org/cancer/types/breast-cancer/about/types-of-breast-cancer/inflammatory-breast-cancer.html</u>

References

American Cancer Society. *Cancer Facts & Figures 2024*. Atlanta: American Cancer Society; 2024.

Young JL Jr, Roffers SD, Ries LAG, Fritz AG, Hurlbut AA (eds). SEER Summary

Staging Manual - 2000: Codes and Coding Instructions, National Cancer Institute, NIH Pub. No. 01-4969, Bethesda, MD, 2001.

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Questions to Ask Your Doctor About Breast Cancer

It's important to be able to have frank, open discussions with your cancer care team. They want to answer all of your questions so that you can make informed treatment and life decisions. Here are some questions that you can use to help better understand your cancer and your treatment options.

- When you're told you have breast cancer
- When deciding on a treatment plan
- If you need surgery
- During treatment
- After treatment
- Preparing your list of questions

When you're told you have breast cancer

- Exactly what type of breast cancer¹ do I have?
- How big is the cancer? Where exactly is it?
- Has the cancer spread to my lymph nodes or other organs?
- What is the stage of my cancer? What does it mean?
- Will I need any other tests before we can decide on treatment?
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- Should I think about <u>genetic testing</u>²? What are my testing options? Should I take a home-based genetic test? What would be the reasons for and against testing?
- How do I get a copy of my pathology report?
- If I'm worried about the costs and <u>insurance coverage</u>³ for my diagnosis and treatment, who can help me?

When deciding on a treatment plan

- How much experience do you have treating this type of cancer?
- Should I get a <u>second opinion</u>⁴? How do I do that? Will getting a second opinion delay my treatment and can that affect my outcome?
- What are my <u>treatment</u>⁵ choices?
- What treatment do you recommend and why?
- Should I think about taking part in a <u>clinical trial</u>⁶?
- What would the goal of the treatment be?
- How soon do I need to start treatment?
- How long will treatment last? What will it be like? Where will it be done?
- Should my biopsy tissue be sent for a gene expression test to help decide if chemotherapy might be helpful for me?
- Are there othermolecular or protein tests that need to be done on my cancer tissue to help decide my treatment options?
- What should I do to get ready for treatment?
- What risks or side effects are there to the treatments you suggest? Are there things I can do to reduce these side effects?
- How will treatment affect my daily activities? Can I still work fulltime?
- Will I lose my hair? If so, what can I do about it?
- Will I go through menopause as a result of the treatment? Will I be able to have children after treatment? Would I be able to breastfeed?
- Do I have time to freeze my eggs before starting treatment? What are my options?
- What are the chances the cancer will come back (recur) after this treatment?
- What would we do if the treatment doesn't work or if the cancer comes back?
- What if I have transportation problems getting to and from treatment?

If you need surgery

- Is breast-conserving surgery (lumpectomy) an option for me? Why or why not?
- What are the positive and negative sides of breast-conserving surgery versus mastectomy?
- How many surgeries like mine have you done?
- Will you have to take out lymph nodes? If so, would you advise a sentinel lymph node biopsy? Why or why not?
- What side effects might lymph node removal cause?
- How long will I be in the hospital?
- Will I have stitches or staples at the surgery site? Will there be a drain (tube) coming out of the site?
- How do I care for the surgery site? Will I need someone to help me?
- What will my breasts look and feel like after my surgery? Will I have normal feeling in them?
- What will the scar look like?
- Is <u>breast reconstruction surgery</u>⁸ an option if I want it? What would it mean in my case?
- Can I have reconstruction at the same time as the surgery to remove the cancer?
 What are the reasons for and against having it done right away or waiting until later?
- What types of reconstruction might be options for me?
- Could you recommend a plastic surgeon I could speak to about reconstruction options?
- Will I need a breast form (prosthesis), and if so, where can I get one?
- Do I need to stop taking any medications or supplements before surgery?
- When should I call your office if I'm having side effects or concerns?

During treatment

Once treatment begins, you'll need to know what to expect and what to look for. Not all of these questions may apply to you, but asking the ones that do may be helpful.

- How will we know if the treatment is working?
- Is there anything I can do to help manage side effects?
- What symptoms or side effects should I tell you about right away?
- How can I reach you on nights, holidays, or weekends?
- Will I need to change what I eat during treatment?
- Are there any limits on what I can do?

health care professionals, such as nurses and social workers, can answer some of your questions.

To find out more about speaking with your health care team, see <u>The Doctor-Patient</u> Relationship¹¹.

Hyperlinks

- 1. www.cancer.org/cancer/types/breast-cancer/about/types-of-breast-cancer.html
- 2. www.cancer.org/cancer/risk-prevention/genetics.html
- 3. <u>www.cancer.org/cancer/financial-insurance-matters/understanding-health-insurance.html</u>
- 4. <u>www.cancer.org/cancer/managing-cancer/finding-care/seeking-a-second-opinion.html</u>
- 5. www.cancer.org/cancer/types/breast-cancer/treatment.html
- 6. <u>www.cancer.org/cancer/managing-cancer/making-treatment-decisions/clinical-trials.html</u>
- 7. <u>www.cancer.org/cancer/types/breast-cancer/treatment/surgery-for-breast-cancer/breast-conserving-surgery-lumpectomy.html</u>
- 8. <u>www.cancer.org/cancer/types/breast-cancer/reconstruction-surgery.html</u>
- 9. www.cancer.org/cancer/managing-cancer/side-effects/swelling/lymphedema.html
- 10. <u>www.cancer.org/cancer/managing-cancer/making-treatment-decisions/clinical-trials.html</u>
- 11. <u>www.cancer.org/cancer/managing-cancer/finding-care/the-doctor-patient-relationship.html</u>

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Our team is made up of doctors and oncology certified nurses with deep knowledge of cancer care as well as editors and translators with extensive experience in medical writing.

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