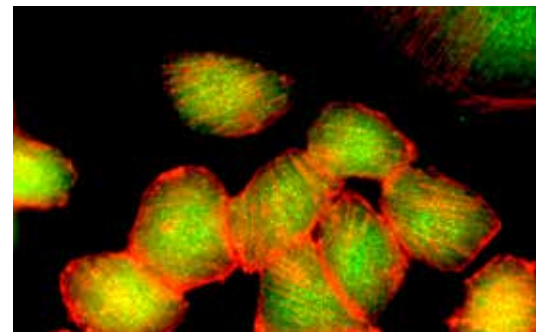
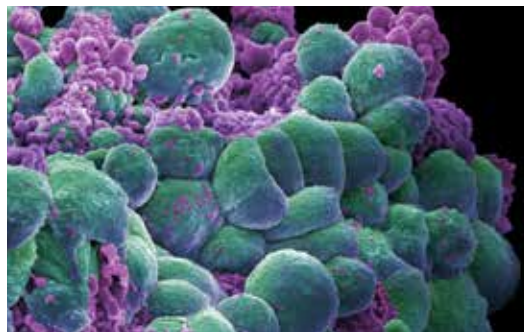
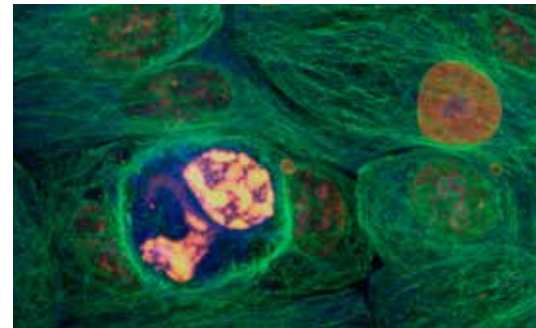
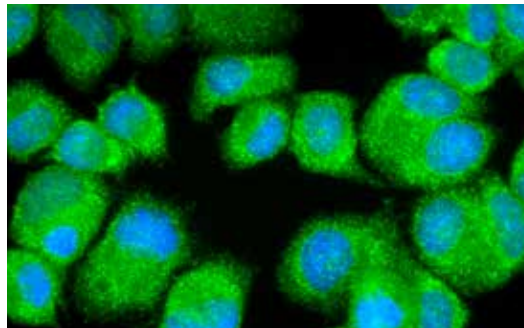


THE ESSENTIALS OF
LIFE SCIENCE RESEARCH
GLOBALLY DELIVERED™

Breast Cancer Resource Book



BREAST CANCER RESOURCE BOOK

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**THE ESSENTIALS
OF LIFE SCIENCE
RESEARCH
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DELIVERED™**

ATCC provides research and development tools and reagents as well as related biological material management services, consistent with its mission: to acquire, authenticate, preserve, develop, and distribute standard reference microorganisms, cell lines, and related materials for research in the life sciences.

For over 85 years, ATCC has been a leading provider of high-quality biological materials and standards to the life science community. We are an independent, 501(c)(3) non-profit entity focused on scientific enablement at universities, research institutes, government agencies, and commercial research labs. Our diverse and comprehensive resources in cell biology and microbiology have been central to the growth of the biotechnology age. ATCC has as its core mission to source,

authenticate and further develop products and services essential to the needs of basic and applied life science work.

ATCC distributes to more than 165 countries on 6 continents and has a growing international network of 12 distribution partners. Our infrastructure and experience in biological materials logistics enables us to work effectively with researchers no matter where they are located.

INTRODUCTION

Worldwide, breast cancer accounts for approximately 23% of all cancer diagnoses in woman, and represents the leading cause of cancer deaths among all female cancer patients¹. Thus, breast cancer affects a large portion of the global population and constitutes a substantial public health burden. Accordingly, it has generated a considerable amount of research interest.

Breast cancers encompass a heterogeneous array of tumor types that are classified according to their histological and molecular characteristics. Classification methods are being refined as molecular profiling techniques improve, but currently breast cancers are sorted into one of at least four subtypes and each subtype is associated with a different prognosis and course of treatment (see Table below). Therefore, to generate the most effective treatment options, investigators need in vitro research tools that represent the heterogeneity of breast cancers in vivo.

This guide will describe the extensive collection of cell lines, primary cells, and associated reagents that ATCC offers to support breast cancer research. We created this guide to assist researchers in the study of this complicated disease. We hope you will find it a helpful resource for planning and getting your experiments up and running..



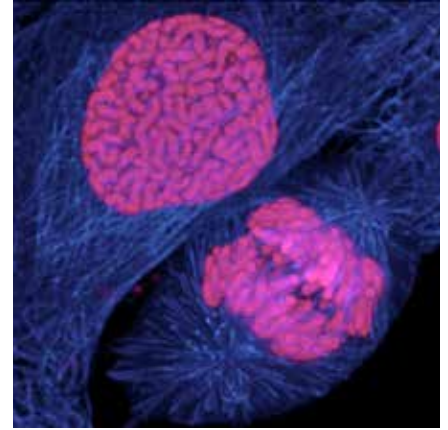
Classifications Breast Cancer Classifications

| Subtype | Immunoprofile | Characteristics ^{2,3} |
|-------------------------|-------------------|--|
| Luminal A | ER+, PR+/-, HER2- | Low expression of proliferation marker Ki67 Responsive to hormone therapy Often responsive to chemotherapy |
| Luminal B | ER+, PR+/-, HER2+ | High expression of proliferation marker Ki67 Usually responsive to hormone therapy Variably responsive to chemotherapy Variably responsive to HER2 antibody therapies (i.e. trastusumab) |
| Basal (Triple Negative) | ER-, PR-, HER2- | High expression of proliferation marker Ki67 Expression of EGFR+ Variable expression of basal cell marker cytokeratin 5/6 Not responsive to hormone therapy, but often responsive to chemotherapy |
| HER2 amplified | ER-, PR-, HER2+ | High expression of proliferation marker Ki67 Often responsive to HER2 antibody therapies (i.e. trastusumab) and chemotherapy |

TUMOR CELL LINES

Tumor cell lines have formed the cornerstone of breast cancer research since the early 1970s, when the MCF-7 line and the MD Anderson series were first established². Since that time, the number of tumor cell lines available to breast cancer researchers has exploded and their contribution to our understanding of this complex disease cannot be overstated. In fact, some of the most effective treatment options available today exist because researchers were able to tease out the disease mechanism, using an in vitro culture system, and design targeted therapeutics.

Tumor cell lines have become even more powerful with the advent of next-generation sequencing technology. The availability of this technology has led to the formation of several large-scale sequencing initiatives, and these have generated a vast amount of actionable data. One such initiative is the Cancer Cell Line Encyclopedia (CCLE). The CCLE is a collaborative effort between Novartis and the Broad Institute, which has released mutation data for 1,651 genes representing nearly 1,000 cell lines. The CCLE research group used this data set to compare the copy number, expression pattern, and mutation frequency of tumor cell lines with primary tumors and showed that tumor cell lines are representative of their in vivo counterparts. Additionally, they used the sequencing data to predict that tumor cell lines harboring particular mutations are sensitive to specific classes of drugs⁴.



Thus, tumor cell lines are an invaluable resource not only for understanding disease mechanisms, but also for drug design and discovery. The following pages contain a description of the ways our cell lines are organized. Please see the appendix for the full listing of the ATCC breast cancer cell lines and for lists of cell lines that have been organized according to gene mutation or that are part of a paired tumor/normal set.

Breast Cancer Cell Lines by Gene

Please see the appendix for a list of the Breast Cancer Cell lines arranged by Gene mutation.

The cell lines in these lists are arranged based on gene mutation information obtained from the Catalogue of Somatic Mutations in Cancer (Wellcome Trust Sanger Institute, UK).

- **CDKN2A** - Cyclin-dependent kinase inhibitor 2A (CDKN2A) is a tumor suppressor gene that encodes at least three different splice variants, two of which can induce arrest at the G1 phase of the cell cycle by inhibiting the CDK4 kinase. Germ-line mutations in this gene are associated with the development of several types of malignancy, including breast cancer⁵.
- **PIK3CA** - The phosphatidylinositol-4, 5-bisphosphate 3-kinase, catalytic subunit alpha (PIK3CA) gene encodes the p110 α catalytic subunit of class I phosphatidylinositol 3-kinases (PI3K). PI3K operates as part of the PI3K/AKT/mTOR pathway to mediate cell proliferation, survival, migration and vesicular trafficking. Mutations of this gene are frequently associated with breast cancer and are considered a positive prognostic factor^{6,7}.
- **PTEN** - Phosphatase and tensin homolog (PTEN) is a tumor suppressor gene that encodes for a lipid phosphatase⁸. PTEN primarily exerts its anti-tumorigenic effect by inhibiting the activation of PI3K. However, PTEN may also help regulate genomic stability, cell cycle progression, differentiation and gene expression⁹. Mutations in this gene are commonly associated with a variety of cancers, including prostate, brain and breast⁸.
- **TP53** - The TP53 gene encodes the tumor suppressor protein p53, which plays a significant role in regulating the cellular response to DNA damage and other cytotoxic stresses. In addition, it plays an important regulatory role in cellular functions including cell cycle arrest, DNA repair, genome stability, apoptosis, cell differentiation and angiogenesis. Mutations in TP53 are commonly associated with human malignancies, and can be found in a significant proportion of human cancers from an array of tissue sources, including breast¹⁰.

Paired Tumor/Normal Cell Lines

Tumor-derived cell lines matched to either normal or metastatic cell lines obtained from the same patient provide a valuable resource for cancer studies. The availability of such models allows researchers to compare tumor lines to their normal or near normal counterparts.

TUMOR CELL PANELS

The value of individual tumor cell lines is further enhanced by organizing them into groups or “panels” according to their histological or molecular characteristics. Such panels can be used for predicting the cellular response of a particular class of disease to novel therapeutics in a controlled, experimental setting.

ATCC has developed an extensive collection of breast-cancer related Tumor Cell Panels to complement our wide array of individual tumor cell lines. Each ATCC Tumor Cell Panel includes low-passage, authenticated tumor cell lines, that have been annotated with genetic mutation data (from the Catalogue of Somatic Mutations in Cancer database, Wellcome Trust Sanger Institute, UK), and collected together in ways that best represent specific features of this heterogeneous disease. The panels are described below to aid you in selecting the one that best suits your research needs.

Please see the appendix for the cell lines included in each Breast Cancer Tumor Cell Panels.

The **Comprehensive Breast Cancer Cell Panel (ATCC® No. 30-4500K™)** is a comprehensive set of 45 breast cancer cell lines derived from ATCC master seed stocks to eliminate variability. Each panel features a CD containing signed certificates of analyses and product sheets for each individual cell line. Please see the website for a full listing of the cell lines included in the panel.

The **Triple Negative Breast Cancer Cell Panels (ATCC® No. TCP-1001, TCP-1002, TCP-1003™)** are arranged according to their classification into the following subtypes: (1) Basal-like, which includes subtypes Basal-Like 1 and 2 (BL1 and BL2) and Immunomodulatory (IM); (2) Mesenchymal-like, which includes the Mesenchymal (M) and Mesenchymal Stem-Like (MSL) groups; and, (3) the Luminal Androgen Receptor (LAR) subtype. **Triple Negative Breast Cancer Panel 1, Basal-Like Morphology** is composed of 10 triple negative breast tumor cell lines that share a basal-like morphology. **Triple Negative Breast Cancer Panel 2, Mesenchymal and Luminal Morphology** is composed of six triple negative breast tumor cell lines that share a mesenchymal-like morphology or a LAR subtype. **Triple Negative Breast Cancer Panel 3**, is composed of all the items in (ATCC® No. TCP-1001) and (ATCC® No. TCP-1002), plus two triple negative breast cancer cell lines with an unclassified morphology.

The **Breast Cancer Biomarkers Cell Panel (ATCC® No. TCP-1004™)** takes the Tumor Cell Panel concept to the next level by including published biomarker data for each culture in a convenient, printable format. This panel puts biomarker information at the researcher’s fingertips, so they can reach a deeper understanding of the mechanisms behind the development and progression of breast cancer.

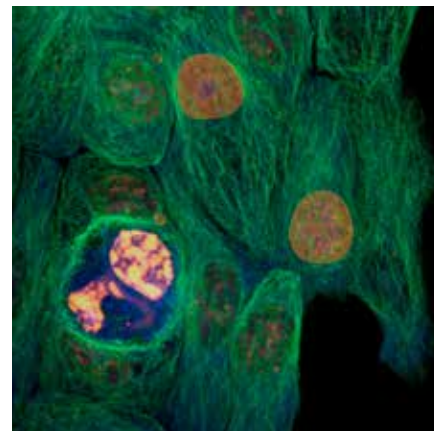
The **Breast Cancer p53 Hotspot Mutation Cell Panel (ATCC® No. TCP-2010™)** is designed to help investigators unravel the relationship between TP53 (p53) gene mutations and oncogenesis. ATCC sequenced the p53-mutant cell lines in our collection and arranged them based on their precise mutational profiles. This panel combines cell lines that harbor mutations at different p53 hotspots with appropriate control lines that are either wild-type or null for p53 expression. Additionally, the p53-mutant cell lines included in these panels have mutations that result in translated proteins that are either unable to bind DNA, or that are structurally altered. Thus, this panel allows researchers to perform mechanistic assays at both the gene and protein level.

The **Breast Cancer Mouse Model Cell Panel (ATCC® No. TCP-1005™)** is composed of eight immortalized mouse mammary epithelial cell lines that stably overexpress MEK1 activated mutant (MEKDD), EGFR2/Neu, Myc or Ha-Ras. The cell lines in this panel have been used successfully to generate mouse models of breast cancer for studying metastasis and EGFR-MEK signaling, oncogenes in cell transformation, and for testing anti-cancer compounds¹¹⁻¹⁴.

Cell authentication

The value of tumor cell lines continues to grow as large-scale, cell-line sequencing initiatives release massive amount of genomic information that can be used to annotate and associate them with a particular class of tumor. However, the immeasurable potential of annotated tumor cell lines is undermined by the ease with which they can become corrupted by intra- or interspecies contamination.

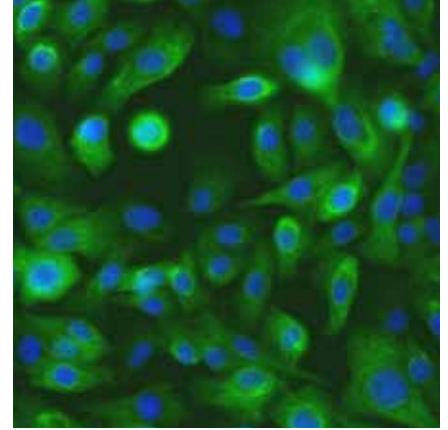
All ATCC tumor cell lines are rigorously tested to confirm their identity and to rule out both intra- and interspecies contamination. These tests include morphology, karyotyping, cytochrome C oxidase I (COI) gene analysis (a PCR-based assay to detect inter-species contamination), and short tandem repeat (STR) profiling.



hTERT IMMORTALIZED CELLS

ATCC offers a wide variety of cells that have been immortalized through the forced expression of the hTERT component of the Telomerase gene. Expression of hTERT allows human cells to maintain the telomere ends of chromosomes and repress replicative senescence. Analysis of numerous hTERT-immortalized cell lines indicates these cells have a stable karyotype and retain many of the physiological characteristics of the primary cell, including normal phenotypic marker expression. Additionally, they exhibit normal p53 cell cycle checkpoint control, are non-malignant, contact inhibited, and anchorage dependent. Moreover, they retain normal growth responses to serum and mitogens, require growth factors for proliferation, and do not show changes associated with transformation, such as tumorigenicity or growth in soft agar¹⁵.

The **human mammary epithelium hTERT-HME1 [ME16C] (ATCC® No.CRL-4010™)** cell line was derived from normal primary mammary epithelial cells. These cells were infected with a retroviral pBabepuro+hTERT vector and cultured in complete growth medium containing puromycin until stable clones were selected¹⁶. hTERT-HME1 cells have served as normal controls in several studies that sought to unravel the molecular mechanism of breast cancer pathogenesis. For example, Zhang and colleagues used this cell line to show that the BRMS1 gene is expressed in normal cells, but not in metastatic cancer cells. They went on to show that this gene sensitizes breast cancer cells to ATP-induced growth inhibition and apoptosis¹⁷. In another study, Lee and colleagues used hTERT-HME1 cells to show that certain Caveolin-1 mutations contribute to the pathogenesis of breast cancer by acting in a dominant-negative manner. Thus, hTERT-HME1 cells have contributed to mechanistic studies that have helped drive basic research and that have the potential to inform rational therapeutic design¹⁸.



ATCC® No. CRL-4010™ stained with a monoclonal pan-cytokeratin antibody (green) and Hoechst dye (blue)

Spectrum of Tools for Cell-Based Research

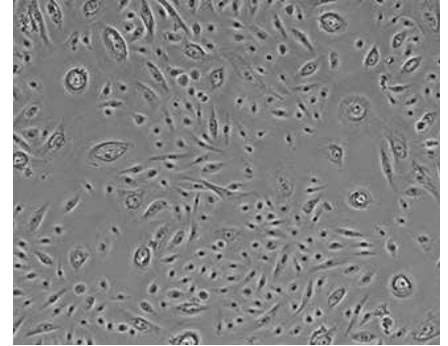
| Cell Type | Tissue Phenotype | Genotypic Stability | Proliferative Capacity | Ease-of-Use |
|-------------------------------|------------------|---------------------|------------------------|-------------|
| Primary Cell Solution | ++++ | ++++ | + | ++ |
| hTERT Immortalized Cell Lines | +++ | +++ | ++++ | ++++ |
| Continuous Cell Lines | ++ | + | ++++ | ++++ |

PRIMARY CELLS

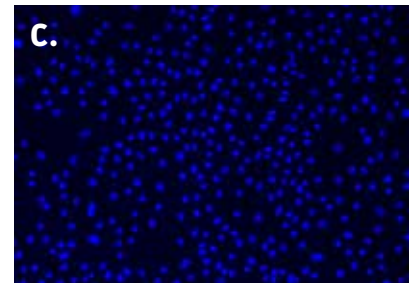
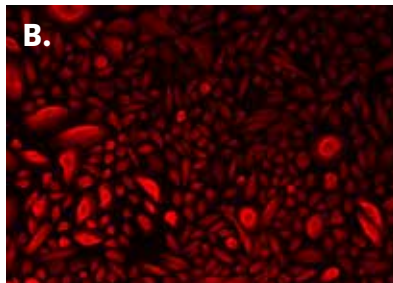
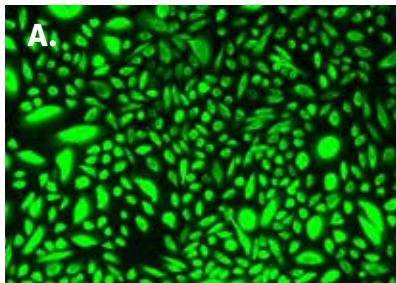
The human mammary gland is a complex tissue composed of milk-producing luminal epithelial cells surrounded by contractile myoepithelial cells; and the vast majority of breast cancers originate in these structures. Studies have suggested that continuous cell lines tend towards lineage-restricted profiles that fail to mimic the cellular heterogeneity of either the normal or cancerous mammary gland¹⁹. Thus, cultures derived directly from the tissue may provide a more representative model for certain applications like the study of oncogenesis or drug discovery.

Primary cells can be isolated in the laboratory setting, but the isolation process makes primary cell cultures vulnerable to contamination by bacteria or non-epithelial cells. Plus, access to an ethically-derived source of normal human mammary tissue may be difficult to obtain. These issues may be avoided entirely by using Primary Human Mammary Epithelial Cells from ATCC.

ATCC **Primary Mammary Epithelial Cells (ATCC® No. PCS-600-010)** are a mixed population of myoepithelial and luminal epithelial cells. They are cryopreserved at low passage (P2) to ensure high post-thaw viability and plating efficiency and they have been thoroughly tested to confirm their proliferative capacity and that they are free of microbial contamination. Together with ATCC Mammary Epithelial Cell Basal Medium and Growth kit they form a complete culture system that you can trust to help you achieve your research objectives.



ATCC® No. PCS-600-010™ Normal, Human Primary Mammary Epithelial Cells



Primary Mammary epithelial cells are a mixed population of CK14-positive myoepithelial cells (A) and CK18-positive luminal epithelial cells (B), DAPI (C).

CULTURE AND ASSAY REAGENTS

ATCC has the media you need to keep your cells healthy and behaving as expected. All ATCC media are filtered, ready to use, and shipped in 500 mL plastic bottles. Our collection includes all the “classic” media formulations, plus media and reagents specially designed to support the culture of mammary cells *in vitro*.

Phenol-red Free Media

Phenol red, which is commonly used as a pH indicator in culture media, bears a structural resemblance to non-steroidal estrogens. Additionally, it has been found to stimulate cell proliferation, in a dose dependent, estrogen receptor-dependent manner²⁰. Consequently, phenol red may mask or exaggerate the impact of experimentally applied estrogen, effectively altering experimental results in unforeseeable ways. To help researchers establish more controllable experimental conditions, ATCC now offers phenol-red free DMEM and RPMI.

Every ATCC media product is manufactured to the exact specifications recommended by ATCC cell culture scientists, and rigorously tested to meet the quality and performance standards you’ve come to expect from ATCC.

- DMEM without phenol red (ATCC® No. 30-2601)
- RPMI without phenol red (ATCC® No. 30-2602)

Primary Mammary Epithelial Cell Complete Media

The Mammary Epithelial Cell Basal Medium and Growth kit work together to provide a complete, serum-free growth media designed to support the proliferation and plating efficiency of epithelial cells derived from normal human breast.

Primary Mammary Epithelial Cell Basal Media (ATCC® No. PCS-600-030) is a sterile, phenol red-free, liquid tissue culture medium that contains essential and non-essential amino acids, vitamins, other organic compounds, trace minerals and inorganic salts.

Primary Mammary Epithelial Cell Growth Kit (ATCC® No. PCS-600-040) includes rH-Insulin, L-Glutamine, Epinephrine, Apo-Transferrin, rH-TGF α , Extract P, and Hydrocortisone Hemisuccinate.

CellMatrix Basement Membrane Gel

The extracellular matrix is important for normal maintenance of the breast epithelium and may be involved in oncogenic transformation². In the *in vitro* setting, CellMatrix gel is used to support a differentiated phenotype in primary epithelial cultures, and it may be used to promote spheroid formation in cancer cell lines. This later feature of CellMatrix may be particularly useful for the screening of anti-cancer drugs. Drugs that are highly effective on cells grown in monolayer cultures are often ineffective when tested in pre-clinical animal models. This is likely due to the fact that cells grown *in vitro* in a monolayer are more accessible to a chemical compound than cells growing *in vivo*, in three-dimensions, within an extracellular matrix. Therefore, CellMatrix Basement Membrane Gel is essential for helping investigators recapitulate the *in vivo* conditions of normal and cancerous cells in an *in vitro* culture model.

CellMatrix Basement Membrane Gel (ATCC® No. ACS-3035) is a soluble, growth factor-reduced basement membrane extract purified from the Engelbreth Holm Swarm (EHS) tumor. It comprises a mixture of laminin, collagen IV, entactin and heparin sulfate proteoglycan, and is commonly used to generate physiologically relevant two and three-dimensional culture conditions. It is supplied as a 12 to 18 mg/mL solution (refer to the product label for the actual concentration) stored in Dulbecco’s Modified Eagle’s Medium (DMEM) with 10 μ g/mL gentamycin, and it has been tested to ensure that it is free from microbial contamination (i.e. bacterial (including mycoplasma), fungus, and 31 other pathogens or viruses, including LDEV, as demonstrated by PCR. Endotoxin concentrations are less than 8 EU/mL by LAL assay).



MTT and XTT Proliferation Assay Kits

The reduction of tetrazolium salts to an easy-to-measure, colored precipitate is widely accepted as a reliable way to examine cell viability and proliferation. ATCC offers proliferation kits that rely on the reduction of the popular tetrazolium dyes MTT or XTT.

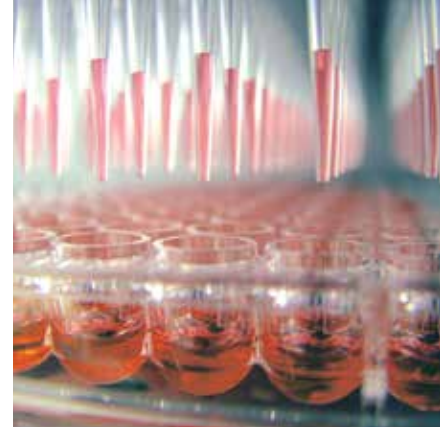
- **MTT Proliferation kit (ATCC® No. 30-1010K)**

The yellow tetrazolium MTT (3-(4, 5-dimethylthiazolyl-2)-2, 5-diphenyltetrazolium bromide) is reduced by metabolically active cells, in part by the action of dehydrogenase enzymes, to generate reducing equivalents such as NADH and NADPH. The resulting intracellular formazan derivative can be solubilized and quantified using a spectrophotometer.

The MTT Cell Proliferation Assay measures the cell proliferation rate and conversely, when metabolic events lead to apoptosis or necrosis, the reduction in cell viability. The number of assay steps has been minimized as much as possible to expedite sample processing. The MTT Reagent yields low background absorbance values in the absence of cells. For each cell type the linear relationship between cell number and signal produced is established, thus allowing an accurate quantification of changes in the rate of cell proliferation.

- **XTT Proliferation Kit (ATCC® No. 30-1011K)**

The second generation tetrazolium dye, XTT, can be effectively used in cell proliferation, cytotoxicity, and apoptosis assays^{21, 22}. XTT is reduced, to a soluble brightly-colored orange derivative, by a mix of cellular effectors. The sensitivity of an XTT assay is greatly improved by using the included intermediate electron carrier, PMS (N-methyl dibenzopyrazine methyl sulfate), to help drive XTT reduction and the formation of the formazan derivative.



Growth Curves

Cells grow at different rates in each of the different phases of the growth cycle and the calculated doubling time may be a composite of growth during more than one of these phases. Growth during exponential growth or log phase is fairly constant and reproducible for a given set of growth conditions.

APPENDIX

Tumor Cell Lines

Complete List of ATCC Breast Cancer Cell Lines

| Tumor Source | Pathology | Organism | Name | ATCC® No. |
|--|------------------|----------|-----------------|-------------|
| Primary | adenocarcinoma | Human | Hs 281.T | CRL-7227™ |
| Primary | adenocarcinoma | Human | Hs 343.T | CRL-7245™ |
| Primary | adenocarcinoma | Human | Hs 362.T | CRL-7253™ |
| Primary | adenocarcinoma | Human | MDA-MB-468 | HTB-132™ |
| Metastasis: brain | adenocarcinoma | Human | MDA-MB-361 | HTB-27™ |
| Metastasis: malignant pleural effusion | adenocarcinoma | Human | AU565 [AU-565] | CRL-2351™ |
| Metastasis: pleural effusion | adenocarcinoma | Human | MCF7 | HTB-22™ |
| Metastasis: pleural effusion | adenocarcinoma | Human | MDA-MB-231 | HTB-26™ |
| Metastasis: pleural effusion | adenocarcinoma | Human | MDA-MB-415 | HTB-128™ |
| Metastasis: pleural effusion | adenocarcinoma | Human | MDA-MB-436 | HTB-130™ |
| Metastasis: pleural effusion | adenocarcinoma | Human | CAMA-1 | HTB-21™ |
| Metastasis: pleural effusion | adenocarcinoma | Human | SK-BR-3 | HTB-30™ |
| Metastasis: pleural effusion | adenocarcinoma | Human | MDA-MB-231 | CRM-HTB-26™ |
| Primary | adenocarcinoma | Mouse | JC | CRL-2116™ |
| Primary | adenocarcinoma | Rat | 13762 MAT B III | CRL-1666™ |
| Primary | adenocarcinoma | Rat | NMU | CRL-1743™ |
| Primary | adenocarcinoma | Rat | RBA | CRL-1747™ |
| Primary | adenocarcinoma | Human | UACC-1179 | CRL-3127™ |
| Metastasis: pleural effusion | adenocarcinoma | Human | UACC-2087 | CRL-3180™ |
| Primary | adenocarcinoma | Human | Hs 274.T | CRL-7222™ |
| Primary | adenocarcinoma | Human | Hs 739.T | CRL-7477™ |
| Primary | adenocarcinoma | Rat | Hs 741.T | CRL-7480™ |
| Primary | adenocarcinoma | Human | SMT/2A LNM | CRL-6602™ |
| Primary | cancer | Human | Hs 748.T | CRL-7486™ |
| Primary | carcinoma | Human | Hs 578Bst | HTB-125™ |
| Primary | carcinoma | Human | Hs 578T | HTB-126™ |
| Primary | carcinoma | Human | BT-20 | HTB-19™ |
| Primary | carcinoma | Human | Hs 579.Mg | CRL-7347™ |
| Primary | ductal carcinoma | Human | UACC-812 | CRL-1897™ |
| Primary | ductal carcinoma | Human | BT-483 | HTB-121™ |
| Primary | ductal carcinoma | Human | BT-549 | HTB-122™ |
| Primary | ductal carcinoma | Human | Hs 574.T | CRL-7345™ |
| Primary | ductal carcinoma | Human | BT-474 | HTB-20™ |
| Metastasis: ascites | ductal carcinoma | Human | ZR-75-1 | CRL-1500™ |
| Metastasis: pleural effusion | ductal carcinoma | Human | T47D-KBluc | CRL-2865™ |
| Metastasis: pleural effusion | ductal carcinoma | Human | T-47D | HTB-133™ |

Please see www.atcc.org for more information about each of the cell lines listed here

Complete List of ATCC Breast Cancer Cell Lines

| Tumor Source | Pathology | Organism | Name | ATCC® No. |
|----------------------------------|--------------------------------------|----------|------------------------|------------|
| Metastasis: pleural effusion | ductal carcinoma | Human | MDA-MB-134-VI | HTB-23™ |
| Metastasis: pleural effusion | ductal carcinoma | Human | MDA-MB-175-VII | HTB-25™ |
| Metastasis: ascites | ductal carcinoma | Human | ZR-75-30 | CRL-1504™ |
| Primary | ductal carcinoma | Human | UACC-2648 | CRL-3121™ |
| Primary | fibrocystic disease | Human | MCF 10A | CRL-10317™ |
| Primary | fibrocystic disease | Human | MCF 10F | CRL-10318™ |
| Primary | fibrocystic disease | Human | MCF 10-2A | CRL-10781™ |
| Primary | infiltrating ductal carcinoma | Human | Hs 564(E).Mg | CRL-7329™ |
| Metastasis: axillary lymph node | infiltrating ductal carcinoma | Human | UACC-3199 | CRL-2983™ |
| Primary | infiltrating ductal carcinoma | Human | Hs 319.T | CRL-7236™ |
| Metastasis: pleural effusion | infiltrating lobular carcinoma | Human | UACC-3133 | CRL-2988™ |
| Metastasis: pleural fluid | inflammatory carcinoma | Human | UACC-732 | CRL-3166™ |
| Primary | carcinoma | Mouse | EMT6 | CRL-2755™ |
| Primary | medullary carcinoma | Human | MDA-MB-157 | HTB-24™ |
| Metastasis: pericardial effusion | carcinoma | Human | MDA-MB-453 | HTB-131™ |
| Metastasis: pericardial effusion | carcinoma | Human | MDA-kb2 | CRL-2713™ |
| Metastasis: pericardial effusion | papilloma | Mouse | CSMalpha6C [CSMab6C] | CRL-8400™ |
| Primary | papilloma | Mouse | CMH1a | CRL-8399™ |
| Primary | papilloma | Mouse | CMalpha1h [CMab1h] | CRL-8401™ |
| Primary | ductal carcinoma | Human | UACC-893 | CRL-1902™ |
| Primary | scirrhous adenocarcinoma | Human | Hs 742.T | CRL-7482™ |
| Primary | cancer | Mouse | 4T1 | CRL-2539™ |
| Primary | ductal carcinoma | Human | HCC1395 | CRL-2324™ |
| Primary | ductal carcinoma | Human | HCC1954 | CRL-2338™ |
| Primary | ductal carcinoma | Human | HCC1008 | CRL-2320™ |
| Primary | ductal carcinoma | Human | HCC1143 | CRL-2321™ |
| Primary | ductal carcinoma | Human | HCC1187 | CRL-2322™ |
| Primary | acantholytic squamous cell carcinoma | Human | HCC1806 | CRL-2335™ |
| Primary | ductal carcinoma | Human | HCC1500 | CRL-2329™ |
| Primary | ductal carcinoma | Human | HCC38 | CRL-2314™ |
| Primary | ductal carcinoma | Human | HCC1937 | CRL-2336™ |
| Primary | ductal carcinoma | Human | HCC202 | CRL-2316™ |
| Primary | ductal carcinoma | Human | HCC1599 | CRL-2331™ |
| Primary | ductal carcinoma | Human | HCC1419 | CRL-2326™ |
| Primary | ductal carcinoma | Human | HCC2218 | CRL-2343™ |
| Primary | ductal carcinoma | Human | HCC70 | CRL-2315™ |
| Primary | metaplastic carcinoma | Human | HCC1569 | CRL-2330™ |
| Metastasis: pleural effusion | adenocarcinoma | Human | HCC1428 | CRL-2327™ |
| Primary | tumor | Mouse | C127I | CRL-1616™ |
| None | normal; spontaneously immortalized | Human | MCF-12A | CRL-10782™ |
| None | normal; fibrocystic disease | Human | MCF-12F | CRL-10783™ |
| Primary | cancer | Human | MDA-kb2 | CRL-2713™ |
| Primary | cancer | Human | Hs 605.T | CRL-7365™* |
| Primary | Paget's disease | Human | SW527 [SW 527, SW-527] | CRL-7940™* |
| Primary | cancer | Human | DU4475 | HTB-123™ |
| None | normal: chemically immortalized | Human | 184A1 | CRL-8798™ |
| None | normal: chemically immortalized | Human | 184B5 | CRL-8799™ |
| Metastasis: pleural effusion | carcinoma | Human | MB157 | CRL-7721™ |

Complete List of ATCC Breast Cancer Cell Lines (continued)

| Tumor Source | Pathology | Organism | Name | ATCC® No. |
|--------------|-------------------------------|----------|-------------|------------|
| Primary | cancer | Monkey | CMMT | CRL-6299™ |
| Primary | cancer | Mouse | MMT 060562 | CCL-51™ |
| Primary | hyperplastic alveolar nodules | Mouse | CL-S1 | CRL-1615™ |
| None | normal | Mouse | NMuMG | CRL-1636™ |
| None | normal | Mouse | MM3MG | CRL-6376™* |
| Primary | tumor | Mouse | MM5MTC | CRL-6378™* |
| Primary | cancer | Mouse | +/+ MGT | CRL-6468™* |
| Primary | tumor | Rat | LA7 | CRL-2283™ |
| Primary | tumor | Mouse | Rn2T | CRL-6599™* |
| None | normal | Dog | CF37.Mg | CRL-6230™* |
| Primary | cancer | Dog | CF41.Mg | CRL-6232™* |
| Primary | cancer | Human | Hs 566(B).T | CRL-7336™* |
| Primary | cancer | Human | Hs 606.T | CRL-7368™* |
| Primary | tumor | Mouse | Mm5MT | CRL-1637™* |
| Primary | cancer | Human | Hs 329.T | CRL-7242™* |
| Primary | cancer | Human | Hs 371.T | CRL-7256™* |
| Primary | cancer | Human | Hs 190.T | CRL-7145™* |
| Primary | cancer | Human | Hs 344.T | CRL-7246™* |
| Primary | cancer | Human | Hs 350.T | CRL-7248™* |
| None | normal | Human | Hs 617.Mg | CRL-7379™* |
| Primary | cancer | Human | Hs 841.T | CRL-7574™* |
| Primary | cancer | Human | Hs 849.T | CRL-7583™* |
| Primary | cancer | Human | Hs 851.T | CRL-7584™* |
| Primary | cancer | Human | Hs 861.T | CRL-7596™* |
| Primary | abnormal | Human | Hs 875.T | CRL-7612™* |
| Primary | cancer | Mouse | MM2MT | CRL-6373™* |
| Primary | cancer | Mouse | RlllMT | CRL-6449™* |
| Primary | cancer | Rat | Rn1T | CRL-6598™* |
| None | normal | Dog | CF38.Mg | CRL-6231™* |
| Primary | tumor | Dog | CF34.Mg | CRL-6228™* |

*Part of the NBL Cell Line Collection. This cell line is neither produced nor fully characterized by ATCC. We do not guarantee that it will maintain a specific morphology, purity, or any other property upon passage.

Breast Cancer Cell Lines by Gene

CDKN2A

| Tumor Source | Pathology | Zygosity | Gene Sequence [†] | Protein Sequence [†] | Name | ATCC® No. |
|------------------------------|--------------------------------------|------------|----------------------------|-------------------------------|------------|-----------|
| Primary | carcinoma | homozygous | c.1471del471 | p.0? | BT-20 | HTB-19™ |
| Primary | acantholytic squamous cell carcinoma | homozygous | c.1471del471 | p.0? | HCC1806 | CRL-2335™ |
| Primary | ductal carcinoma | homozygous | c.1471del471 | p.0? | HCC38 | CRL-2314™ |
| Primary | ductal carcinoma | homozygous | c.1471del471 | p.0? | HCC1395 | CRL-2324™ |
| Metastasis; pleural effusion | adenocarcinoma | homozygous | c.1471del471 | p.0? | MCF7 | HTB-22™ |
| Metastasis; pleural effusion | adenocarcinoma | homozygous | c.1471del471 | p.0? | MDA-MB-231 | HTB-26™ |
| Metastasis; brain | adenocarcinoma | homozygous | c.156G>C | p.M521 | MDA-MB-361 | HTB-27™ |

[†]For a description of the sequence variation nomenclature please refer to: *den Dunnen JT and Antonarakis SE (2000), Hum. Mutat. 15:7-12.*

PIK3CA

| Tumor Source | Pathology | Zygoty | Gene Sequence [†] | Protein Sequence [†] | Name | ATCC [®] No. |
|---------------------------------|------------------|--------------|----------------------------|-------------------------------|------------|-----------------------|
| Primary | carcinoma | heterozygous | c.1616C>G | p.P539R | BT-20 | HTB-19™ |
| Primary | carcinoma | heterozygous | c.3140A>G | p.H1047R | BT-20 | HTB-19™ |
| Primary | ductal carcinoma | heterozygous | c.3140A>G | p.H1047R | HCC1954 | CRL-2338™ |
| Primary | ductal carcinoma | heterozygous | c.3140A>G | p.H1047R | UACC-893 | CRL-1902™ |
| Primary | ductal carcinoma | heterozygous | c.333G>C | p.K111N | BT-474 | HTB-20™ |
| Metastasis; pleural effusion | adenocarcinoma | heterozygous | c.1633G>A | p.E545K | MDA-MB-361 | HTB-22™ |
| Metastasis; pleural effusion | carcinoma | heterozygous | c.3140A>G | p.H1047R | MCF7 | HTB-131™ |
| Metastasis; pleural effusion | ductal carcinoma | heterozygous | c.3140A>G | p.H1047R | MDA-MB-453 | HTB-133™ |
| Metastasis; brain | adenocarcinoma | heterozygous | c.1633G>A | p.E545K | T-47D | HTB-27™ |

PTEN

| Tumor Source | Pathology | Zygoty | Gene Sequence [†] | Protein Sequence [†] | Name | ATCC [®] No. |
|---------------------------------|----------------------------|--------------|----------------------------|-------------------------------|------------|-----------------------|
| Primary | ductal carcinoma | homozygous | c.270delT | p.F90fs*9 | HCC70 | CRL-2315™ |
| Primary | ductal carcinoma | homozygous | c.635_1212del578 | p.N212fs*1 | HCC1395 | CRL-2324™ |
| Primary | papillary ductal carcinoma | homozygous | c.823delG | p.V275fs*1 | BT-549 | HTB-122™ |
| Metastasis; pleural effusion | adenocarcinoma | homozygous | c.253+1G>T | p.? | MDA-MB-468 | HTB-132™ |
| Metastasis; pleural effusion | adenocarcinoma | heterozygous | c.274G>C | p.D92H | CAMA-1 | HTB-21™ |
| Metastasis; pleural effusion | adenocarcinoma | homozygous | c.407G>A | p.C136Y | MDA-MB-415 | HTB-128™ |
| Metastasis; pleural effusion | adenocarcinoma | heterozygous | c.831_834delCTTC | p.T277fs*13 | CAMA-1 | HTB-21™ |

TP53

| Tumor Source | Pathology | Zygoty | Gene Sequence [†] | Protein Sequence [†] | Name | ATCC [®] No. |
|---------------------------------|--------------------------------------|--------------|--|-------------------------------|------------|-----------------------|
| Primary | ductal carcinoma | homozygous | c.1024C>T | p.R342* | UACC-893 | CRL-1902™ |
| Primary | ductal carcinoma | homozygous | c.220_226delGCCCTG | p.A74fs*47 | HCC1419 | CRL-2326™ |
| Primary | ductal carcinoma | homozygous | c.322_324delGGT | p.G108del | HCC1187 | CRL-2322™ |
| Primary | carcinoma | homozygous | c.394A>C | p.K132Q | BT-20 | HTB-19™ |
| Primary | ductal carcinoma | homozygous | c.488A>G | p.Y163C | HCC1954 | CRL-2338™ |
| Primary | ductal carcinoma | homozygous | c.524G>A | p.R175H | HCC1395 | CRL-2324™ |
| Primary | ductal carcinoma | homozygous | c.659A>G | p.Y220C | HCC1419 | CRL-2326™ |
| Primary | ductal carcinoma | homozygous | c.673-2A>T | p.? | HCC1599 | CRL-2331™ |
| Primary | ductal carcinoma | homozygous | c.743G>A | p.R248Q | HCC70 | CRL-2315™ |
| Primary | ductal carcinoma | homozygous | c.743G>A | p.R248Q | HCC1143 | CRL-2321™ |
| Primary | papillary ductal carcinoma | homozygous | c.747G>C | p.R249S | BT-549 | HTB-122™ |
| Primary | acantholytic squamous cell carcinoma | homozygous | c.766_767insAA | p.T256fs*90 | HCC1806 | CRL-2335™ |
| Primary | ductal carcinoma | homozygous | c.818G>T | p.R273L | HCC38 | CRL-2314™ |
| Primary | ductal carcinoma | homozygous | c.847C>T | p.R283C | HCC2218 | CRL-2343™ |
| Primary | ductal carcinoma | homozygous | c.853G>A | p.E285K | BT-474 | HTB-20™ |
| Primary | metaplastic carcinoma | heterozygous | c.880G>T | p.E294* | HCC1569 | CRL-2330™ |
| Primary | ductal carcinoma | homozygous | c.916C>T | p.R306* | HCC1937 | CRL-2336™ |
| Metastasis; pleural effusion | medullary carcinoma | homozygous | c.261_286delAGCCCCCTCTGGCCCCCTGTCATCTT | p.A88fs*52 | MDA-MB-157 | HTB-24™ |

[†]For a description of the sequence variation nomenclature please refer to: *den Dunnen JT and Antonarakis SE (2000), Hum. Mutat. 15:7-12.*

TP53 (continued)

| Tumor Source | Pathology | Zygoty | Gene Sequence [†] | Protein Sequence [†] | Name | ATCC [®] No. |
|------------------------------|----------------|------------|----------------------------|-------------------------------|------------|-----------------------|
| Metastasis; pleural effusion | adenocarcinoma | homozygous | c.524G>A | p.R175H | AU565 | CRL-2351™ |
| Metastasis; pleural effusion | adenocarcinoma | homozygous | c.580C>T | p.L194F | T-47D | HTB-133™ |
| Metastasis; pleural effusion | adenocarcinoma | homozygous | c.707A>G | p.Y236C | MDA-MB-415 | HTB-128™ |
| Metastasis; pleural effusion | adenocarcinoma | homozygous | c.818G>A | p.R273H | MDA-MB-468 | HTB-132™ |
| Metastasis; pleural effusion | adenocarcinoma | homozygous | c.839G>A | p.R280K | MDA-MB-231 | HTB-26™ |
| Metastasis; pleural effusion | adenocarcinoma | homozygous | c.839G>C | p.R280T | CAMA-1 | HTB-21™ |

Other genes

| Tumor Source | Pathology | Gene | Zygoty | Gene Sequence [†] | Protein Sequence [†] | Name | ATCC [®] No. |
|------------------------------|----------------------------|--------|--------------|----------------------------|-------------------------------|------------|-----------------------|
| Metastasis; skin | ductal carcinoma | APC | homozygous | c.4729G>T | p.E1577* | DU4475 | HTB-123™ |
| Metastasis; skin | ductal carcinoma | BRAF | heterozygous | c.1799T>A | p.V600E | DU4475 | HTB-123™ |
| Metastasis; pleural effusion | adenocarcinoma | BRAF | heterozygous | c.1391G>T | p.G464V | MDA-MB-231 | HTB-26™ |
| Metastasis; ascites | ductal carcinoma | PIK3R1 | homozygous | c.335_427del93 | p.? | ZR-75-30 | CRL-1504™ |
| Metastasis; pleural effusion | adenocarcinoma | RAS | heterozygous | c.38G>A | p.G13D | MDA-MB-231 | HTB-26™ |
| Primary | papillary ductal carcinoma | RB1 | homozygous | c.265_607del343 | p.? | BT-549 | HTB-122™ |
| Metastasis; skin | ductal carcinoma | RB1 | homozygous | c.265_2787del2787 | p.0? | CAMA-1 | HTB-21™ |
| Metastasis; pleural effusion | adenocarcinoma | RB1 | homozygous | c.265_2787del2523 | p.? | MDA-MB-468 | HTB-132™ |
| Metastasis; pleural effusion | adenocarcinoma | SMAD4 | Homozygous | c.1_1659del1659 | p.0? | MDA-MB-468 | HTB-132™ |

Paired Tumor/Normal Cell Lines

| Tumor Cell Lines Tumor Source | Pathology | Name | ATCC [®] No. | Normal Pairing Tissue Source | Pathology | Name | ATCC [®] No. |
|-------------------------------|------------------|----------|-----------------------|------------------------------|-----------|------------|-----------------------|
| Metastasis: lymph node | ductal carcinoma | HCC10008 | CRL-2320™ | B lymphoblast | normal | HCC1007 BL | CRL-2319™ |
| Mammary gland | ductal carcinoma | Hs574.T | CRL-7345™ | Skin | normal | Hs574.Sk | CRL-7346™ |
| Mammary gland | ductal carcinoma | Hs578T | HTB-126™ | Mammary gland | normal | Hs578Bst | HTB-125™ |
| Mammary gland | ductal carcinoma | HCC1954 | CRL-2338™ | B lymphoblast | normal | HCC1954 BL | CRL-2339™ |
| Mammary gland | ductal carcinoma | HCC38 | CRL-2314™ | B lymphoblast | normal | HCC38 BL | CRL-2346™ |
| Mammary gland | ductal carcinoma | HCC1143 | CRL-2321™ | B lymphoblast | normal | HCC1143 BL | CRL-2362™ |
| Mammary gland | ductal carcinoma | HCC1187 | CRL-2322™ | B lymphoblast | normal | HCC1187 BL | CRL-2323™ |
| Mammary gland | ductal carcinoma | HCC1395 | CRL-2324™ | B lymphoblast | normal | HCC1395 BL | CRL-2325™ |
| Mammary gland | ductal carcinoma | HCC1599 | CRL-2331™ | B lymphoblast | normal | HCC1599 BL | CRL-2332™ |
| Mammary gland | ductal carcinoma | HCC1937 | CRL-2336™ | B lymphoblast | normal | HCC1937 BL | CRL-2337™ |
| Mammary gland | ductal carcinoma | HCC2218 | CRL-2343™ | B lymphoblast | normal | HCC2218 BL | CRL-2363™ |

[†]For a description of the sequence variation nomenclature please refer to: *den Dunnen JT and Antonarakis SE (2000), Hum. Mutat. 15:7-12.*

Tumor Cell Panels

Comprehensive Breast Cancer Cell Panel (ATCC® No. 30-4500K™)

| Tumor Source | Pathology | Organism | Name | ATCC® No. |
|--|--------------------------------------|----------|----------------|------------|
| None | normal: chemically-transformed | Human | 184B5 | CRL-8799™ |
| Metastasis: malignant pleural effusion | adenocarcinoma | Human | AU565 [AU-565] | CRL-2351™ |
| Primary | carcinoma | Human | BT-20 | HTB-19™ |
| Primary | ductal carcinoma | Human | BT-474 | HTB-20™ |
| Primary | ductal carcinoma | Human | BT-483 | HTB-121™ |
| Primary | ductal carcinoma | Human | BT-549 | HTB-122™ |
| Metastasis: pleural effusion | adenocarcinoma | Human | CAMA-1 | HTB-21™ |
| Primary | fibrocystic disease | Human | MCF 10A | CRL-10317™ |
| Primary | fibrocystic disease | Human | MCF 10F | CRL-10318™ |
| Primary | fibrocystic disease | Human | MCF-10-2A | CRL-10781™ |
| Primary | infiltrating ductal carcinoma | Human | Hs 564(E).Mg | CRL-7329™ |
| Metastasis: axillary lymph node | infiltrating ductal carcinoma | Human | UACC-3199 | CRL-2983™ |
| Metastasis: pleural effusion | infiltrating lobular carcinoma | Human | Hs 319.T | CRL-7236™ |
| Metastasis: pleural effusion | inflammatory carcinoma | Human | UACC-3133 | CRL-2988™ |
| Primary | carcinoma | Mouse | UACC-732 | CRL-3166™ |
| Primary | carcinoma | Mouse | HCC1500 | CRL-2329™ |
| Metastasis: pericardial effusion | medullary carcinoma | Human | HCC1569 | CRL-2330™ |
| Primary | ductal carcinoma | Human | HCC1599 | CRL-2331™ |
| Primary | acantholytic squamous cell carcinoma | Human | HCC1806 | CRL-2335™ |
| Primary | ductal carcinoma | Human | HCC1937 | CRL-2336™ |
| Primary | ductal carcinoma | Human | HCC1954 | CRL-2338™ |
| Primary | ductal carcinoma | Human | HCC2218 | CRL-2343™ |
| Primary | carcinoma | Human | Hs 578Bst | HTB-125™ |
| Primary | carcinoma | Human | Hs 578T | HTB-126™ |
| Metastasis: pleural effusion | adenocarcinoma | Human | MCF7 | HTB-22™ |
| Primary | fibrocystic disease | Human | MCF 10A | CRL-10317™ |
| Primary | fibrocystic disease | Human | MCF 10F | CRL-10318™ |
| None | normal; spontaneously immortalized | Human | MCF-12F | CRL-10782™ |
| Primary | Cancer | Human | MDA-kb2 | CRL-2713™ |
| Metastasis: pleural effusion | ductal carcinoma | Human | MDA-MB-134-VI | HTB-23™ |
| Primary | medullary carcinoma | Human | MDA-MB-157 | HTB-24™ |
| Metastasis: pleural effusion | ductal carcinoma | Human | MDA-MB-175-VII | HTB-25™ |
| Metastasis: pleural effusion | adenocarcinoma | Human | MDA-MB-231 | HTB-26™ |
| Metastasis: brain | adenocarcinoma | Human | MDA-MB-361 | HTB-27™ |
| Metastasis: pleural effusion | adenocarcinoma | Human | MDA-MB-415 | HTB-128™ |
| Metastasis: pleural effusion | adenocarcinoma | Human | MDA-MB-436 | HTB-130™ |
| Metastasis: pericardial effusion | carcinoma | Human | MDA-MB-453 | HTB-131™ |
| Primary | adenocarcinoma | Human | MDA-MB-468 | HTB-132™ |
| Metastasis: pleural effusion | adenocarcinoma | Human | SK-BR-3 | HTB-30™ |
| Metastasis: pleural effusion | ductal carcinoma | Human | T-47D | HTB-133™ |
| Primary | ductal carcinoma | Human | UACC-812 | CRL-1897™ |
| Primary | ductal carcinoma | Human | UACC-893 | CRL-1902™ |
| Metastasis: ascites | ductal carcinoma | Human | ZR-75-1 | CRL-1500™ |
| Metastasis: ascites | ductal carcinoma | Human | ZR-75-30 | CRL-1504™ |

Triple Negative Breast Cancer Cell Panel 1 (ATCC® No. TCP-1001™)

| Tumor Source | Pathology | Subtype* | Mutant Gene | Zygoty | Gene Sequence† | Protein Sequence† | Name | ATCC® No. |
|---------------------------------|--|----------|----------------------------------|--|---|-------------------------------------|------------|-----------|
| Primary | Primary ductal carcinoma | BL1 | BRCA2 TP53 | homozygous homozygous | c.4550_4559del10 c.673-2A>T | p.K1517fs*23 p.? | HCC1599 | CRL-2331™ |
| Primary | Primary ductal carcinoma | BL1 | BRCA TP53 | homozygous homozygous | c.5266_5267insC c.916C>T | p.Q1756fs*74 p.R306 | HCC1937 | CRL-2336™ |
| Primary | Primary ductal carcinoma | BL1 | TP53 | homozygous | c.743G>A | p.R248Q | HCC1143 | CRL-2321™ |
| Metastasis; pleural effusion | Adenocarcinoma | BL1 | PTEN RB1 SMAD4 TP53 | homozygous homozygous homozygous homozygous | c.253+1G>T c.265_2787del2523 c.1_1659del1659 c.818G>A | p.? p.? p.0? p.R273H | MDA-MB-468 | HTB-132™ |
| Primary | Primary ductal carcinoma | BL1 | CDKN2A TP53 | homozygous homozygous | c.1_471del471 c.818G>T | p.0? p.R273L | HCC38 | CRL-2314™ |
| Primary | Primary ductal carcinoma | BL2 | PTEN TP53 | homozygous homozygous | c.270delT c.743G>A | p.F90fs*9 p.R248Q | HCC70 | CRL-2315™ |
| Primary | Primary acantholytic squamous cell carcinoma | IM | CDKN2A KDM6A STK11 TP53 | homozygous homozygous homozygous homozygous | c.1_471del471 c.444_564del121 c.1109_1302del194 c.766_767insAA | p.0? p.0 p.? p.T256fs*90 | HCC1806 | CRL-2335™ |
| Primary | Primary ductal carcinoma | IM | TP53 | homozygous | c.322_324delGGT | p.G108del | HCC1187 | CRL-2322™ |
| Metastasis; skin | Carcinoma | IM | APC BRAF MAP2K4 RB1 | homozygous heterozygous homozygous homozygous | c.4729G>T c.1799T>A c.1_1200del1200 c.1_2787del2787 | p.E1577* p.V600E p.0? p.0? | DU4475 | HTB-123™ |

Triple Negative Breast Cancer Cell Panel 2 (ATCC® No. TCP-1002™)

| Tumor Source | Pathology | Subtype* | Mutant Gene | Zygoty | Gene Sequence† | Protein Sequence† | Name | ATCC® No. |
|---------------------------------|---------------------|----------|---------------------------------------|--|---|---|------------|-----------|
| Primary | Ductal carcinoma | M | PTEN RB1 TP53 | homozygous homozygous homozygous | c.823delG c.265_607del343 c.747G>C | p.V275fs*1 p.? p.R249S | BT-549 | HTB-122™ |
| Primary | Carcinoma | MSL | CDKN2A HRAS PIK3R1 TP53 | homozygous heterozygous homozygous homozygous | c.1_471del471 c.35G>A c.1358_1359insTAA c.469G>T | p.0? p.G12D p.N453_454insN p.V157F | Hs578T | HTB-126™ |
| Metastasis; pleural effusion | Adenocarcinoma | MSL | BRAF CDKN2A KRAS NF2 TP53 | heterozygous homozygous heterozygous homozygous homozygous | c.1391G>T c.1_471del471 c.38G>A c.691G>T c.839G>A | p.G464V p.0? p.G13D p.E231* p.R280K | MDA-MB-231 | HTB-26™ |
| Metastasis; pleural effusion | Adenocarcinoma | MSL | BRCA1 RB1 | homozygous homozygous | c.5277+1G>A c.607_608ins227 | p.? p.G203fs*9 | MDA-MB-436 | HTB-130™ |
| Metastasis; pleural effusion | Medullary carcinoma | MSL | NF1 TP53 | homozygous homozygous | c.8253_8268del16 c.261_286delAGCCCCCTCGGCCCTGTCATCTT | p.S2751fs*27 p.A88fs*52 | MDA-MB-157 | HTB-24™ |
| Metastasis; pleural effusion | Carcinoma | LAR | CDH1 PIK3CA | homozygous heterozygous | c.1913G>A c.3140A>G | p.W638* p.H1047R | MDA-MB-453 | HTB-131™ |

*For a description of the sequence variation nomenclature please refer to: *den Dunnen JT and Antonarakis SE (2000), Hum. Mutat. 15:7-12.*

Triple Negative Breast Cancer Cell Panel 3 (ATCC® No. TCP-1003™): Cell lines in Panels 1 and 2 above plus the two unclassified cell lines listed below

| Tumor Source | Pathology | Subtype* | Mutant Gene | Zygoty | Gene Sequence† | Protein Sequence† | Name | ATCC® No. |
|--------------|--------------------------|----------|------------------------------------|--|--|---|---------|-----------|
| Primary | Carcinoma | - | CDKN2A PIK3CA PIK3CA TP53 | homozygous heterozygous heterozygous homozygous | c.1_471del471 c.1616C>G c.3140A>G c.394A>C | p.0? p.P539R p.H1047R p.K132Q | BT-20 | HTB-19™ |
| Primary | Primary ductal carcinoma | - | BRCA1 CDKN2A PTEN TP53 | homozygous homozygous homozygous homozygous | c.5251C>T c.1_471del471 c.635_1212del578 c.524G>A | p.R1751* p.0? p.N212fs*1 p.R175H | HCC1395 | CRL-2324™ |

*These subtypes are classified as: (1) Basal-like, including subtypes BL1 (basal-like 1), BL2 (basal-like 2) and IM (immunomodulatory); (2) Mesenchymal-like, including subtypes M (mesenchymal) and MSL (mesenchymal stem-like); and, (3) LAR (luminal androgen receptor).

Breast Cancer Biomarkers Cell Panel (ATCC® No. TCP-1004™)

| Tumor Source | Pathology | Age | Positive Markers | Negative Markers | Other Significant Features | Patient Treatment | Name | ATCC® No. |
|------------------------------|-------------------------------|-----|------------------|--|---|--|-----------|-----------|
| Primary | infiltrating ductal carcinoma | 43 | HER-2/neu | ER, PR, EGFR, P-glycoprotein | - | Vinblastine, Adriamycin, Cytosin, Cyclophosphamide, Methotrexate, 5-fluorouracil | UACC-812 | CRL-1897™ |
| Primary | infiltrating ductal carcinoma | 57 | HER-2/neu | ER, PR, EGFR, P-glycoprotein, MASPIN | MASPIN promoter methylation has been reported for this line | None | UACC-893 | CRL-1902™ |
| Metastasis; axillary nodes | infiltrating ductal carcinoma | 58 | EGFR | ER, PR, HER-2/ Neu | Methylated BRCA-1 promoter | Cytosin, Adriamycin, 5-fluorouracil, Tamoxifen, Mitoxantrone, Vinblastine | UACC-3199 | CRL-2983™ |
| Metastasis; pleural effusion | ductal carcinoma | 63 | HER-2/neu, BMP-3 | ER (very low), PR, EGFR, MASPIN, DSC3, BMP-2 | MASPIN promoter methylation has been reported for this line | Surgery only | UACC-3133 | CRL-2988™ |
| Metastasis; pleural effusion | adenocarcinoma | 62 | HER-2/neu | ER, PR, EGFR, MASPIN, DSC3 | P53 R213X mutation and MASPIN promoter methylation have been reported for this line. | Adriamycin, Cytosin, Methotrexate, Tamoxifen | UACC-1179 | CRL-3127™ |
| Metastasis; pleural effusion | adenocarcinoma | 35 | HER-2/neu, PR | ER, EGFR | Drug resistant cell line to cyclin D kinase 4/6 inhibitor and HER-2 inhibitors. | Vinblastine, Adriamycin, Cytosin | UACC-732 | CRL-3166™ |
| Metastasis; pleural effusion | adenocarcinoma | 53 | EGFR | ER, PR, HER-2/ Neu, vimentin, MASPIN, DSC3 | P53 V216M mutation has been reported in this cell line. It has also been reported that the MASPIN promoter is not methylated. | Cyclophosphamide, Methotrexate, 5-fluorouracil, Thymidine phosphorylase, Tamoxifen | UACC-2087 | CRL-3180™ |

Breast Cancer Biomarkers Cell Panel (ATCC® No. TCP-1004™)

| Tumor Source | Overexpression | Significant Features | Name | ATCC® No. |
|---------------------|----------------|--|-------------|-----------|
| Mouse mammary gland | Empty vector | The cells were immortalized from mouse mammary epithelial cell line with an empty vector. They are useful as a control. | Eph4Ev | CRL-3063™ |
| Mouse mammary gland | Mutant MEK1 | The cells stable overexpression of glu-glu epitope-tagged MEK constant activated mutant: Asp218/Asp222 MEK1 phosphorylation site mutant (MEKDD). | B-MEKDD 116 | CRL-3069™ |
| Mouse mammary gland | Mutant MEK1 | The cells were derived from a mouse primary breast tumor. The cells stably overexpress glu-glu epitope-tagged MEK1 constitutively activated mutant (MEKDD). | Eph4 1424 | CRL-3071™ |
| Mouse mammary gland | Mutant MEK1 | The cells were derived from a mouse breast tumor metastasis to lung. The cells stably overexpress glu-glu epitope-tagged MEK1 constitutively activated mutant (MEKDD). | Eph4 1424.1 | CRL-3209™ |
| Mouse mammary gland | Mutant MEK1 | The cells were derived from a mouse breast tumor metastasis to kidney. The cells stably overexpress glu-glu epitope tagged MEK1 constitutively activated mutant (MEKDD). | Eph4 1424.2 | CRL-3210™ |
| Mouse mammary gland | Myc | The cells are a mouse mammary tumor cell line derived from c-myc-initiated transgenic mice. | M158 | CRL-3086™ |

The mutation data was obtained from the Sanger Institute Catalogue Of Somatic Mutations In Cancer web site, <http://www.sanger.ac.uk/cosmic> Bamford et al (2004) The COSMIC (Catalogue of Somatic Mutations in Cancer) database and website. Br J Cancer, 91,355-358. ATCC and The Sanger Institute provide these data in good faith, but make no warranty, express or implied, nor assumes any legal liability or responsibility for any purpose for which the data are used.

†For a description of the sequence variation nomenclature please refer to: *den Dunnen JT and Antonarakis SE (2000), Hum. Mutat. 15:7-12.*

Breast Cancer Biomarkers Cell Panel (ATCC® No. TCP-1004™)

| Tumor Source | Overexpression | Significant Features | Name | ATCC® No. |
|---------------------|----------------|--|--------|-----------|
| Mouse mammary gland | EGFR/Neu | The cells are a mouse mammary tumor cell line derived from Neu-initiated transgenic mice. | NF639 | CRL-3090™ |
| Mouse mammary gland | Ha-Ras | The cells are a mouse mammary tumor cell line derived from Ha-ras-initiated transgenic mice. | Ac 711 | CRL-3092™ |

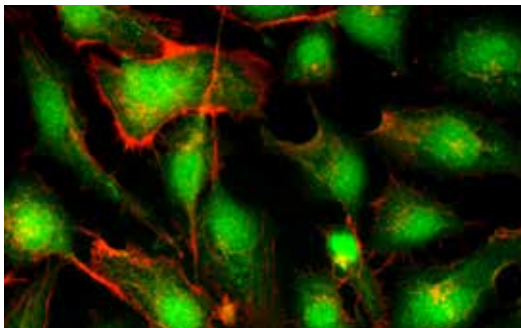
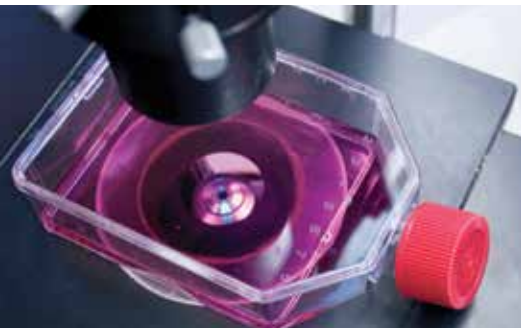
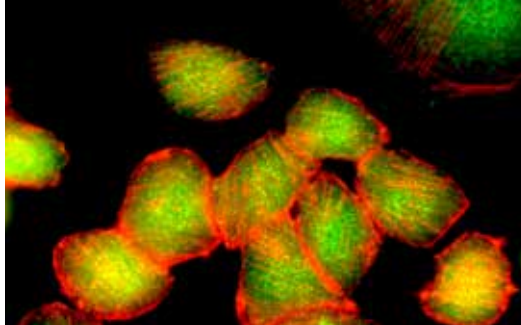
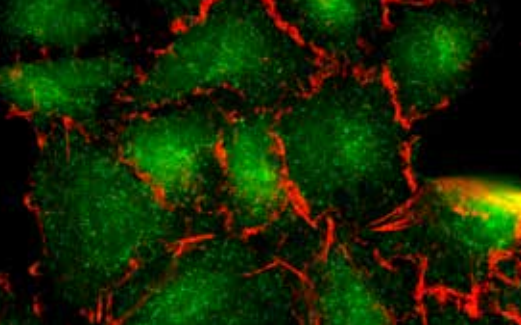
Breast Cancer p53 Hotspot Mutation Cell Panel (ATCC Cat. No. TCP-2010)

| Tumor Source | Pathology | p53 Status | Zygoty | Gene Sequence† | Protein Sequence† | Name | ATCC® No. |
|------------------------------|------------------|------------|------------|----------------|-------------------|----------------|-----------|
| Primary | ductal carcinoma | MUT | homozygous | c.743G>A | p.R248Q | HCC70 | CRL-2315™ |
| Primary | ductal carcinoma | MUT | homozygous | c.747G>C | p.R249S | BT-549 | HTB-122™ |
| Primary | ductal carcinoma | MUT | homozygous | c.818G>T | p.R273L | HCC38 | CRL-2314™ |
| Metastasis; pleural effusion | ductal carcinoma | WT | - | - | - | MDA-MB-175-VII | HTB-25™ |
| Metastasis; pleural effusion | adenocarcinoma | MUT | homozygous | c.524G>A | p.R175H | AU565 | CRL-2351™ |
| Metastasis; pleural effusion | adenocarcinoma | MUT | homozygous | c.524G>A | p.R175H | SK-BR-3 | HTB-30™ |
| Metastasis; pleural effusion | adenocarcinoma | MUT | homozygous | c.818G>A | p.R273H | MDA-MB-468 | HTB-132™ |
| Metastasis; brain | adenocarcinoma | WT | - | - | - | MDA-MB-361 | HTB-27 |

†For a description of the sequence variation nomenclature please refer to: *den Dunnen JT and Antonarakis SE (2000), Hum. Mutat. 15:7-12.*

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Tel 800.638.6597
703.365.2700
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