An Eye on Breast and Prostate Cancers

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Abstract
Cancer is a class of diseases characterized by out-of-control cell growth. There are over 100 different types of cancer, and each is classified by the type of cell that is initially affected. In all these types of cancers breast and prostate cancers are the most prevalent and dangerous. In this review article a field of vision is given on these two hazardous cancers.

Keywords: Breast cancer; Prostate cancer; Cancer therapy; Biomarkers; Vacuum-assisted breast biopsy; Prostate specific antigen testing

Introduction
Cancer is a major public health problem in the United States and many other parts of the world. Currently, one in four deaths in the United States is due to cancer [1]. Breast cancer and prostate cancer are the two most common invasive cancers in women and men, respectively. Breast cancer is one of the most frequently diagnosed cancers among women, with an estimated 184,450 new cases and 40,930 deaths in 2008 [2]. Prostate cancer (PC) is the second leading cause of cancer deaths in men in America and Western Europe [3]. Prostate cancer is most frequently diagnosed among men, with an estimated 192,280 and 27,360 deaths in 2009 [1]. Although these cancers arise in organs that are different in terms of anatomy and physiological function both organs require gonadal steroids for their development, and tumors that arise from them are typically hormone-dependent and have remarkable underlying biological similarities [4].

Risk factors

Genetic: Recent rapid increases in our understanding of the genetic basis of breast cancer have created a resurgent interest in identifying those at increased risk because of family history. The risk of breast cancer associated with a family history is higher in younger women, some studies suggest that the risk is particularly high in women under 40 and then falls off rapidly to a background level of elevated risk. Nine studies reported the risk of breast cancer in women whose daughter had had the disease [5]. A case-control study was performed to estimate the relative risk of developing prostate cancer for men with a positive family history. Prostate cancer had a higher degree of familiality than either breast or colon cancer two cancers with well-recognized familial tendencies. In a recent study involving 691 prostate cancer cases and the male relatives of their wives, an increased risk of prostate cancer among men was observed whose first or second degree relatives also had prostate cancer [6].

Hormonal: Breast cancer is a steroid hormone–dependent tumor. Many studies have reported a correlation between elevated estrogen blood levels and breast cancer and this observation has raised controversy concerning the long-term use of hormonal replacement therapy [7]. Stratification of patients according to hormone (ER/PR) receptor status and nodal metastasis is of great therapeutic importance [8]. Several years ago, only age, race, and family history were known as risk factors for this diseases. However, today, much progress has been made towards discovering hormonal risk factors for prostate cancer [9]. Prostate cancer is a hormone-dependent, epithelial-derived tumor, resulting from uncontrolled growth of genetically unstable transformed cells [10]. High androgen receptor (AR) level in primary tumor predicts increased prostate cancer-specific mortality. However, the mechanisms that regulate AR function in prostate cancer are poorly known [11].

Dietary
High-energy intake in relation to physical activity, which accelerates growth and the onset of menstruation during childhood, leads to weight gain in middle life and thus can contribute substantially to breast cancer risk. Some evidence suggests that vitamin A or other compounds in vitamin A-rich foods may reduce breast cancer risk modestly, but these findings are not conclusive and deserve further consideration [12]. Breast cancer incidence and mortality rates decrease with environmental conditions that promote Vitamin D synthesis in human skin including lower latitude and higher personal exposure to sunlight [13].

While a number of dietary factors have been linked to prostate cancer the evidence is still tentative [14]. Evidence supports little role for dietary fruits and vegetables in prostate cancer occurrence [15]. Red meat and processed meat also appear to have little effect [16]. Lower blood levels of vitamin D may increase the risk of developing prostate cancer [17]. This may be linked to lower exposure to ultraviolet (UV) light, since UV light exposure can increase vitamin D in the body [18].

Medical exposure
Moderate to high-dose radiotherapy is known to increase the risk of breast cancer. Uncertainties remain about the effects of low-dose chest

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X-rays, particularly in individuals at increased genetic risk [19]. There are also some links between prostate cancer and medications, medical procedures, and medical conditions [20]. Use of the cholesterol-lowering drugs known as the statins may also decrease prostate cancer risk [21].

Viral

Many viruses have been associated with human breast cancers, including Epstein-Barr and Cytomegalovirus. New evidence has revealed the frequent presence of high risk human papillomavirus (HPV) strains HPV16 and HPV18 in breast carcinoma biopsies [22]. In 2006, researchers associated a previously unknown retrovirus, Xenotropic MuLV-related virus or XMRV, with human prostate tumors [23]. Subsequent reports on the virus have been contradictory. A group of US researchers found XMRV protein expression in human prostate tumors [24].

Racial differences

In case of breast cancer the hypothesis that white women with breast cancer survive longer than black women with this disease was evaluated in a retrospective analysis of a population-based prospective follow-up study. The racial difference in survival was greatest among women with advanced disease, and a higher proportion of black women with advanced disease did not receive surgery. Even when the type of surgery and stage of disease were controlled, race was a significant prognostic factor [25].

In case of prostate cancer, black men represent a particularly high risk group for the development of prostate cancer. But little is known of the molecular genetic factors, tumor suppressor genes and/or oncogenes that play a role in prostate cancer in black men. Increased research efforts are needed in order to understand this problem at the molecular level [26].

Metabolic syndrome and breast cancer

Metabolic syndrome components are hypertension, dyslipidemia, glucose intolerance and obesity. Metabolic syndrome, also called insulin resistance syndrome, is defined as a constellation of metabolic abnormalities, including glucose intolerance (fasting plasma concentrations \(_{\text{mg/dL}}\) of 110 mg/dL), dyslipidemia (serum triacylglycerol concentration > 150 mg/dL, serum HDL concentration < 40 mg/dL), hypertension (blood pressure > 130/85 mm Hg), and obesity (a waist circumference of > 102 cm) [27]. In addition to steroid hormones, polypeptide growth hormones (insulin-like factors and insulin itself) are thought to play an important role in regulation of breast tumor proliferation [28].

Hypothesis was raised that breast cancer risk is determined by cell proliferation in response to sex hormones, but also to polypeptide growth hormones. Clinical and epidemiological evidence suggests that both breast cancer and the metabolic disorders included in the insulin resistance syndrome are of polygenic and multifactorial origin. Experimental evidence suggests that hyperinsulinemia and its magnitude of the association is modest.

Diagnosis

The first cancer screening test to be widely used was the Pap test. The test was first developed by George Papanicolaou as a method in understanding the menstrual cycle. He also identified Pap tests
potential for early detection of cervical cancer. In 1960s mammography was developed for identification of breast cancer. Later early detection of cervix, breast, colon, rectum, endometrium, prostate, thyroid, oral cavity, skin, lymph nodes, testes, and ovaries cancers were identified and practiced in the clinic [32].

The earliest breast cancers are detected by a mammogram. Mammography and clinical breast exam are used for screening breast cancer. This indicates an approximate likelihood that the lump is cancer, and may also detect some other lesions [33]. When the tests are inconclusive Fine Needle Aspiration and Cytology (FNAC) may be used. FNAC may be done in a GP’s office using local anaesthetic if required, involves attempting to extract a small portion of fluid from the lump. Clear fluid makes the lump highly unlikely to be cancerous, but bloody fluid may be sent off for inspection under a microscope for cancerous cells. Together, these three tools can be used to diagnose breast cancer with a good degree of accuracy. Other options for biopsy include core biopsy, where a section of the breast lump is removed, and an excisional biopsy, where the entire lump is removed. In addition vacuum-assisted breast biopsy (VAB) may help diagnose breast cancer among patients with a mammographically detected breast in women [34]. A data suggest that c-erbB-2 and p53 expression in breast cancer have prognostic significance [35]. It has been demonstrated that the tumor angiogenic ability is one of the most important predictors of breast cancer progression. Factors controlling tumor angiogenesis are varied and currently under investigation. Chemokines, which produced by both tumor cells and cells of tumor microenvironment are known with a role in tumor angiogenesis [36]. Elastic-scattering spectroscopic techniques are also used for the diagnosis of breast cancer for preliminary clinical results [37].

The last 20 years have seen remarkable changes in both the diagnosis and treatment of prostate cancer. Two driving forces have been the development of prostate-specific antigen (PSA) testing as well as the availability of effective pharmacologic interventions to reduce or eliminate the effects of androgens. In case of prostate cancer, for early diagnosis Prostate Specific Antigen testing (PSA testing) can be done. This test gives a better short term outcomes after diagnosis [38]. Ultrasound (US) and Magnetic Resonance Imaging (MRI) are the two main imaging methods used for prostate cancer detection. If cancer is suspected, a biopsy is offered expediently. During a biopsy a urologist or radiologist obtains tissue samples from the prostate via the rectum. A biopsy gun inserts and removes special hollow-core needles (usually three to six on each side of the prostate) in less than a second. Prostate biopsies are routinely done on an outpatient basis and rarely require hospitalization. Fifty-five percent of men report discomfort during prostate biopsy [39].

Now a day’s molecular biomarkers are playing key role in diagnosis of both breast and prostate cancer. A biomarker can be a substance that is introduced into an organism as a means to examine organ function or other aspects of health. There are different types of biomarkers like

- Clinical biomarkers
- Prognostic biomarkers
- Predictive biomarkers
- Biomarkers for early detection
- Biomarkers based on cancer types

The process of the detection methods had made simple yet somewhat complicated though according to a review [40].

**General mechanism of cancer development**

Cell division is a physiological process that occurs in almost all tissues and under many circumstances. Under normal circumstances, the balance between proliferation and programmed cell death, usually in the form of apoptosis, to maintain the integrity of organs and tissues. Certain mutations in DNA tumor suppressor genes get deactivated, lead to cancer, disrupting the orderly physiological processes (Figure 1).

Mutations that can lead to breast cancer have been experimentally linked to estrogen exposure [41]. These mutations are either inherited or acquired after birth. Normal cells divide as many times as needed and stop. They attach to other cells and stay in a place in tissues. This is benign breast disease [42]. These cells become cancerous when mutations destroy their ability to stop dividing, to attach to other cells and to stay where they belong. A prospective study of benign breast disease and the risk of breast cancer confirm the remarked increase in

![Figure 1: General mechanism of cancer development.](image-url)
breast cancer risk among women with atypical hyperplasia, particularly in premenopausal women, and suggest that these women should be encouraged to undergo frequent breast cancer screening [43]. Presumably, they allow the other mutations, which allow uncontrolled division, lack of attachment, and metastasis to distant organs [44].

Osteopontin is a secreted phosphoprotein which plays a critical role in metastasis of breast cancer. The canonical pathway for osteopontin signaling focuses on its binding interactions with integrin and CD44 cell surface receptors. However, the binding characteristic of osteopontin to integrin and CD44 receptors has not been previously examined and its research work is still going on. These additional receptors may play key functions in OPN’s critical role in cancer metastasis [45].

**Stages of breast cancer**

Breast cancer staging using the TNM system is based on the size of the tumor (T), whether or not the tumor has spread to the lymph nodes (N) in the armpits, and whether the tumor has metastasized (M) (i.e. spread to a more distant part of the body). Larger size, nodal spread, and metastasis have a larger stage number and a worse prognosis.

- **Stage 0** is a pre-cancerous or marker condition, either ductal carcinoma in situ (DCIS) or lobular carcinoma in situ (LCIS).
- **Stages 1–3** are within the breast or regional lymph nodes.
- **Stage 4** is ‘metastatic’ cancer that has a less favorable prognosis.

Prostate cancer is considered as malignant tumor because it is a mass of cells that can invade other parts of the body. This invasion of other organs is called metastasis. Prostate cancer most commonly metastasizes to the bones, lymph nodes. It may also invade rectum, bladder and lower ureters after local progression. The route of metastasis to bone is thought to be venous as the prostatic venous plexus draining the prostate connects with the vertebral veins [46]. Interactions between prostate cancer cells and mature bone-producing osteoblasts resulted in gene expression changes that may increase cells’ ability to establish bone metastasis. The potential for inducing expression changes seems to be crucial dependent of the differentiation status of osteoblasts [47].

Prostate cancer is staged to describe the amount and location of cancer. Staging systems also are used to determine treatment regimens. A “TNM” system is used to stage prostate cancer. It takes into consideration PSA level, tumor size, node involvement and amount of metastatic involvement. The “T” represents the tumor tissue size, the “N” represents the amount of lymph node involvement, and the “M” represents the extent to which the cancer has spread within the pelvic region (metastases). Each letter is combined with a number to explain the amount of variation among levels.

**Treatments**

There are different types of therapies and treatments related to breast and prostate cancers like surgery, radiotherapy, chemotherapy, gene therapy, hormone replacement therapy, oncogene therapy, radical prostatectomy, laparoscopy, minimally invasive treatment, tumor immunotherapy, genetic immunotherapy, radionuclide therapy, targeted cancer treatments, adjuvant therapy, immunotherapy, angiogenic chemotherapy, chemo radiation, conventional radiotherapy etc.

**Surgery**

Surgery involves the physical removal of the tumor, typically along with some of the surrounding tissue and frequently sentinel node biopsy.

- **Standard surgeries for breast cancer include:**
  - Mastectomy: Removal of the whole breast.
  - Quadrantectomy: Removal of one quarter of the breast.
  - Lumpectomy: Removal of a small part of the breast.

If the patient desires, then breast reconstruction surgery, a type of cosmetic surgery, may be performed to create an aesthetic appearance. In other cases, women use breast prostheses to simulate a breast under clothing, or choose a flat chest. Women should be informed that arm problems are expected but non-threatening sequel of initial surgical treatment for breast cancer [48].

Surgical treatment for prostate cancer includes radical prostatectomy. This means surgical removal of all or part of the prostate gland by an incision.

**Chemotherapy**

The use of preoperative neoadjuvant chemotherapy in the treatment of breast cancer has been evolving during the past two decades. The benefits of neoadjuvant chemotherapy are: 1) shrinking large cancer tumors so that they are small enough to be removed by breast-conserving surgery instead of radical mastectomy; 2) reducing the chances of spread through the bloodstream, clearing the tiny metastatic lesion, and lowering the possibility of drug fast; 3) understanding the sensitivity of the tumor to chemotherapy and also providing evidence to post-surgery chemotherapy. In summary it was suggested that 18F-FDG imaging with hybrid PET/CT provides a cost-effective method, which could be well accepted for the low-income population in developing countries, to monitor early tumor response after three cycles of neoadjuvant chemotherapy in stage II and III breast cancer by using the reduction of N/T ratio as predictor [49]. For aggressive tumors, the literature recommends CHOP type chemotherapy and mastectomy with lymph node resection, if needed, for management of PNHBL [50].

The first approved chemotherapy agents, mitoxantrone and estramustine (Emcyt), were utilized for prostate cancer patients to help control pain and to relieve symptoms associated with their metastatic disease. Because of the drugs’ inherent side effects, doctors were reluctant to prescribe chemotherapy unless patients were experiencing problems from the cancer. The next phase of research for chemotherapy is to find ways to make Taxotere-based chemotherapy better. One such promising agent, bevacizumab (Avastin), appears to do just that. Avastin is a monoclonal antibody that targets the VEGF receptor; its main mechanism of action is via anti-angiogenesis. The role of chemotherapy is gaining acceptance for men with high risk or metastatic prostate cancer. Now a day’s interferons are also being targeted for treatment of prostate and breast cancer. Interferons (IFNs) may target cancer cells both through their regulation of the immune response, effect on angiogenesis and through direct effect on cancer cells. Unconjugated interferon induced growth arrest [51].

Resistance to apoptosis leads to the progression of many solid cancers. Azurin, a potent anticancer redox protein secreted by *Pseudomonas aeruginosa* (*P. aeruginosa*) species has been reported to have activity against breast cancer cell lines; *P. aeruginosa* MTCC
Recent advances in treatment of breast and prostate cancer

Recent advances in diagnosis and treatment of cancer have led to an increase in cancer survival and hence, there is a greater emphasis on quality beside quantity of survival. However little is known about the Quality Of Life of survivors [60]. Azurin, a potent anticancer redox protein secreted by Pseudomonas aeruginosa (P. aeruginosa) species has been reported to have activity against breast cancer cell lines; P. aeruginosa MTCC (Microbial Type culture collection) 2453 was the strain that secreted the most azurin and showed remarkable apoptosis in breast carcinoma cells like T-47D and ZR-75-1 [52]. Butyric acid (BA), an extracellular metabolite produced from periodontopathic bacteria result in progression of oral cancer such as oral squamous cell carcinoma (OSCC) [53]. Emerging prognostic factors such as Nottingham prognostic index (NPI) or triple-negative status might improve the models currently used by clinicians [54]. Intensity Modulated Radiation Therapy (IMRT) is widely accepted as an appropriate method to treat tumors at many different anatomic locations including lungs [55]. Chemokines and Chemokine receptors play a role in Prostate Cancer Development and Progression. Bone marrow aspiration along with trephine biopsy is essential for the diagnosis and management of multiple myeloma [56].

Radiotherapy

Radiotherapy is one of the main treatments for breast and prostate cancers. Although treatment with a combination of radiotherapy with chemotherapy, and/or surgery improves the prognosis of patients, no biomarkers can distinguish between the responses obtained with the combined therapies. Therefore studies are with patients treated with radiotherapy alone to evaluate surviving as a predictor for radiotherapy. SWOG trial S0102 showed significant activity of the combination of docetaxel and vinorelbine in HER2-negative metastatic breast cancer (MBC). The combination of trastuzumab, docetaxel, and vinorelbine is effective as first-line chemotheraphy in HER2-positive MBC with minimal toxicity. Postoperative adjuvant chemoradiotherapy was recommended as the standard treatment for patients with rectal cancer because it reduces local recurrence. This paradigm shifted with the use of neoadjuvant chemoradiotherapy, which not only reduces local recurrence but also improves sphincter preservation and surgical outcomes [57].

Radiotherapy is a common treatment for prostate cancer, but failure is observed 30 to 40% of the time. It is more common in patients with abnormal p16. It is more common in patients with abnormal p16. The phytochemical diferuloylmethane (curcumin) a naturally occurring flavinoid derived from the rhizome of Curcuma longa, shows potential radiosensitizing effects. In the present study, the effect of curcumin and radiation on cell viability, apoptosis and clonogenic cell death was examined in LNCaP (wild type p16) and PC3 (mutant p16) prostate cancer cells. The cell viability and cell proliferation decreased in presence of both curcumin and radiation as compared to curcumin or radiation alone in both the cell lines. Regardless of p16 status, combination treatment with curcumin (2.5 to 10 μM) and radiation (2 Gy) had a synergistic effect on clonogenic cell death in both LNCaP and PC3 cells. Curcumin appears to radiosensitize prostate cancer cells and may be a possible adjuvant to radiotherapy in the treatment of cancer. Thus there is a Synergistic Effect between Curcumin (diferuloylmethane) and Radiation on Clonogenic Cell Death Independent of p16 in Prostate Cancer Cells [58].

Many viruses have been associated with human breast cancers, human papilloma virus (HPV) strains HPV16 and HPV18 in breast carcinoma shows different phenotypic changes. HPV16 and HPV18 significantly inhibited proliferation and adhesion of T-47D cells, although viability was not affected. Differential effects on proliferation were observed in MCF7 cells; HPV16 inhibited proliferation, while HPV18 stimulated proliferation. No measurable effects in adhesion or viability in MCF7 cells were observed. The phenotypic changes in T-47D and MCF7 cells were associated with changes in mRNA expression of caspase-2, -3 and -8, but not p16 [59].
by understanding the complexity of this heterogeneous group of tumors. Although triple-negative breast cancers are reported to respond to neoadjuvant chemotherapy, survival of patients with such tumors is still poor and their management may therefore require a more aggressive alternative intervention. Therefore, research is being carried out for a reliable biomarker or a target that could be used to individualize both patient prognosis and therapy is essential for the prevention and cure of triple negative breast cancers [71]. In recent years, many studies have examined the relationship between breast cancer outcomes and vitamin D. Vitamin D also plays an important role in breast cancer symptom management. Based on the evidence of ongoing trials, vitamin D supplementation may eventually be used as a cardioprotective agent in breast cancer patients [72].

In case of Prostate cancer using prostate specific antigen (PSA) as an indicator for the presence of bony metastases rather than routine bone scans would have large economic savings given the population size. However, these recommendations are restricted only to symptomatic screening and cannot extrapolate to population-based screening program due to the sample selection of the study [73]. Multicellular Spheroid Culture (MCS) models have become a mainstream culture model for tumor biology and identification of anticancer resistance mechanisms as an alternative to the classical 2D culture models that poorly reflect the structural characteristics seen in vivo. A review has highlighted the characteristics of prostate cancer MCS (P53 and PARP expression, and epigenetics), and underlined the tumor microenvironment as target for prostate cancer therapy. MCS culture models appear to contribute to the identification of cancer stem cells from the prostate. Further studies are needed to clarify mechanisms such as epigenetic regulation, to better characterize the formation of MCS, and to apply this knowledge into prostate cancer biology and the discovery of new targets for prostate cancer [74]. A nanoparticle based imaging agent that targets SPARC (Secreted Protein Acidic Rich in Cysteine), a molecular marker of prostate cancer metastatic potential has been developed. In vivo imaging information correlates with the metastatic potential of the prostate tumor. This prognostic information could enable doctors to stratify patients and design personalized treatment strategies [75].

A review article on prostate cancer provides information on 4 new potential treatments (cardiac hormones) of prostate cancer and their molecular targets (Ras-MEK 1/2-ERK 1/2 kinase cascade). Vessel dilator, long-acting natriuretic peptide, atrial natriuretic peptide and kaliuretic peptide each significantly decrease up to 97% of human prostate cancer cells in culture. After binding to specific receptors, their signaling in prostate cancer cells involves at 95% inhibition of the activation of Ras, 98% inhibition of the activity of MEK 1/2 kinases, and 96% inhibition of the phosphorylation of ERK 1/2 kinases mediated by cyclic GMP. They also inhibit DNA synthesis within prostate cancer cells mediated by cyclic GMP [76]. Combinations of phytoestrogens with radiation therapy and other antioxidants in advanced or metastatic prostate cancer can be considered because there are limited effective therapy options for this group of patients [77]. A current review states that Prostate Cancer Prevention Trial [PCPT] and Reduction by Dutasteride of Prostate Cancer Events [REDUCE] trials reveal that 5-ARI’s have the ability to reduce low grade prostate cancer, but also show a slight increase high grade prostate cancer diagnosis. Follow-up studies suggest that these findings may be due to detection bias, but official guidelines recommend against 5-ARI use due to the potential risk. Efficacy of chemoprevention agents enables: prevention of over-diagnosis, decreased use of overly aggressive treatment, and enhanced quality of life by reducing suffering, mortality, and economic burdens [78]. A comparative proteomic study of proteins in cultivated cells DU-145, PC-3, LNCaP and BPH-1, as well as in prostate tissue samples obtained from patients with malignant and benign tumors is being done [79].

Cancer treatment involving chemotherapy is typically accompanied by toxic side effects, thereby limiting the amount of the drug that can be given to a patient. As a result, all of the tumor tissue may not be exposed to a lethal dose of the drug. The use of nanocarriers such as liposomes and micelles can improve the pharmacological properties of traditional chemotherapeutics. Encapsulated drug is unable to undergo nanocarrier/endosomal escape. However, a combinatorial approach in which surface modifications made to nanocarriers involve both a targeting ligand as well as a p53-triggered release mechanism following cellular internalization based on endosomal pH may in fact generate a very efficacious construct. This type of combinatorial approach in nanocarrier design may in fact prove to someday be a very efficacious drug delivery system commonly used in cancer therapy [80]. Pre-treatment dose verification is carried out at almost all the centers but measurement techniques and plan acceptance criteria are institution specific. Thus, a variety of IMRT QA program in totality is being followed at the Indian hospitals. There is a need to evolve a national protocol for IMRT QA so that treatment outcomes of all the IMRT centers of country can be compared [81].

The cancer stem cell theory proposes that there is a small but constant subpopulation of cancer cells with stem cell properties responsible for the self renewal capacity and unlimited proliferation of tumor as well as increased resistance to antineoplastic drugs. Targeting these cells might constitute an effective way to cure cancer. All glioma cells have stem cells properties but their phenotype varies depending on the environmental conditions. This model provides an alternative explanation to different and sometimes controversial experimental findings and might be a useful guide for future research in the field of gliomas and stem cell biology [82]. The use of the RC, has the potential to increase the successful rate and maybe, overcome the bottleneck when translating preclinical research into clinical trials. From the economical perspective, predicting which drug will successfully pass clinical trials will have a tremendous impact in the drug industry by lowering cost and time by stopping clinical trials of high number of drugs. On the other hand the use of the RC, will help at early stages to decide which drugs have little chance to be useful for monotherapy but might have use in combinational regimes [83]. MU-Tomo can perform in a few seconds an independent dose calculation accurately and provide a secondary check for a point dose validation of helical tomotherapy plans [84]. To provide more clinically convenient image fusions for adaptive radiotherapy (ART), an automatic rigid and deformable image registration framework (AIRF) is developed for multimodal visualization of multiple chronological images and multiple radiotherapy (RT) plans. The AIRF can automatically register multiple volumetric image datasets of patients taken over an extended period of time and can merge multiple RT plans based on different planning CT images for 4D or adaptive radiotherapy [85]. Incorporation of a pertinent tumor endothelial biomarker into a drug development process could also reduce the time, size and cost of clinical trials through the selection of patients who are responding to novel therapies, as well as increasing the chances of regulatory approval of new leads. Finally, the capability to monitor patient response to a targeted treatment through the read out of the vascular integrity can be used to rescue drugs that may benefit specific populations but whose
clinical efficacy was missed in unselected populations dominated by non-responders [86]. Any discussion on cancer biology, directly or indirectly, includes stem cells, in particular, tumor initiating cells and mesenchymal stem cells (MSCs). The question of targeting cancer stem cells and also the supporting MSCs in cancer cannot be achieved by one entity, but partnership among academia, biotechnology and pharmaceuticals [87]. The field of nanomedicine has proved a feasibility of active targeting using different types of ligand as well as nanoparticles. To this end, we emphasize a need of novel ligand beyond antibodies that targets different tumor component including those in the tumor microenvironment to achieve comprehensive cancer treatment [88].

**Conclusion**

This review gives an overall view on two most prevalent and hazardous cancers, the breast and prostate. It addresses the risk factors like genetical, hormonal, dietary etc. Relation between these cancers and metabolic syndrome is given. Mainly it covers all the recent advances in treatment of breast and prostate cancers. Further research in the treatment of these cancers can yield fruitful results.

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