

REVIEW

Breast cancer in Mongolia: an increasingly important health policy issue

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Medical Image Optimization and Perception Group (MIOPeG), Faculty of Health Science, The University of Sydney, Sydney, NSW, Australia Abstract: Breast cancer is a leading cause of cancer-related death for women in both developed and developing countries. The incidence and mortality of breast cancer in Mongolia, while low compared with other counties, has been increasing on an annual basis. In addition, in Mongolia, approximately 90% of the patients are diagnosed at a late stage, resulting in high mortality, with the majority of individuals diagnosed with breast cancer dying within 5 years of diagnosis. Breast cancer screening plays an important role in reducing mortality in Western countries and has been adopted by a number of Asian countries; however, no such approach exists in Mongolia. In a country of limited resources, implementation of expensive health strategies such as screening requires effective allocations of resources and the identification of the most effective imaging methods. This requirement relies on recent accurate data; however, at this time, there is a paucity of information around breast cancer in Mongolia. Until data around features of the disease are available, effective strategies to diagnose breast cancer that recognize the economic climate in Mongolia cannot be implemented and the impact of breast cancer is likely to increase.

Keywords: incidence, mortality, breast screening, Mongolia

Data source and material

The data used in this review mainly arose from 2 sources; the Cancer Registry, located in the National Cancer Center (NCC) in Mongolia,¹ and the International Association of Cancer Research (IACR).² The Cancer Registry in Mongolia was initiated as a hospital-based registry at the NCC in the early 1960s and has now developed as a population-based registry,³ where data on incidence and mortality of breast cancer between 2009 and 2013 are available. All newly diagnosed cancer and death cases along with the personal and clinical information are recorded at the primary- and secondary-level hospitals and submitted to the Cancer Registry via Health info-2 software.⁴

The quality of the data in Mongolia is not clear, as data collection relies on certain factors such as population coverage and collection method. For example, the limited availability of health facilities and trained health professionals in rural areas may result in a number of cancer cases not being diagnosed and many deaths not being recorded. Also, the comprehensiveness of the data is not well understood since histology and morphology records have only become available since 2012.⁴

Introduction

Breast cancer remains the major health concern since it is the most common cancer among women in the world.^{2,5} The rates in incidence and mortality, however, vary

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significantly by region. In general, developed nations demonstrate a greater incidence (greater than 80 per 100,000) compared with developing countries (less than 30 per 100,000), although the latter group is currently demonstrating more rapid increases in incidence.⁶ Women in low- and middle-income countries tend to be diagnosed at late stages largely due to lack of available routine mammographic screening and breast cancer awareness.^{7,8} In such low resource countries, research data are needed to understand the impact and tailoring of limited resources to maximize impacts on breast cancer.⁹⁻¹¹

Mongolia, with a population of 3 million, is one of the fastest developing countries in Asia, but encounters a substantial and unique cancer burden nationwide. While liver cancer is almost 4 times higher than the world average, being by far the most common cancer among both sexes,³ breast cancer is the 5th most common primary cancer among Mongolian women with a current rate of 6 per 100,000 women, accounting for 6% of all new cases of female cancer.¹ Despite these low figures, the incidence rate appears to be increasing on an annual basis.^{1,12} Also, since the majority of cancer patients are diagnosed at a relatively late stage, the disease has become a serious public health issue.^{13,14}

It should be acknowledged from the outset that in Mongolia, details on the incidence, nature and mortality around breast cancer, along with information on type of women affected are limited and without such data, future health policies or systems around this disease cannot be strategically planned or implemented. This review aims to bring together

the available data and provide an overview of the current status of breast cancer epidemiology in Mongolia. The data will be presented in the context of what is known about the disease in Asian countries and westernized populations.

Breast cancer incidence

While Mongolia has a relatively low breast cancer incidence rate, this value has been consistently increasing. An examination of the annual cancer registry in Mongolia between 2009 and 2013, which contains some of the most recent breast cancer data (Figure 1), demonstrated a substantial increase in the breast cancer incidence rate from 3.0 in 2009 to 6.0 in 2013. However, the year on year increase was inconsistent, with the most rapid increase of 54% seen in the 2009–2010 period resulting in incidence rates of 3.0 and 4.6 per 100,000 in 2009 and 2010, respectively. An increasing trend in breast cancer has also been reported earlier by Troisi et al, 12 based on data between 1998 and 2005 from the IACR and the NCC in Mongolia.

As noted in other Asian countries, incidence rates within Mongolia vary by geographical regions (Figure 2). For example, some rural provinces of Khentii and Bayan-Ulgii have considerably higher rates of 10.1 and 11.0/100,000, respectively, compared with the national level at 6.0/100,000,¹ and it is interesting to note that these provinces are the home for most of the ethnic minorities of Mongolia. Such ethnic-dependent variation in incidence (and presentation) of breast cancer has been reported in the neighboring country of People's Republic of China. ¹⁵ Although the cancer registry is

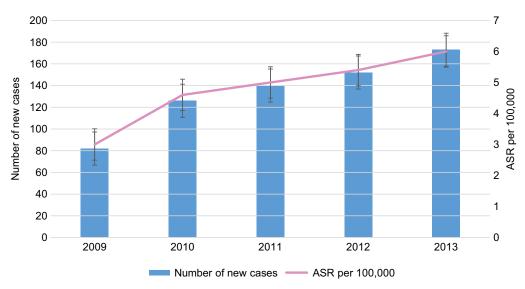


Figure I Incidence rate of female breast cancer, Mongolia, 2009–2013.

Note: Data from Cancer Registry Data, National Cancer Center, Mongolia, 2009–2013.

Abbreviation: ASR, age-standardized rate.

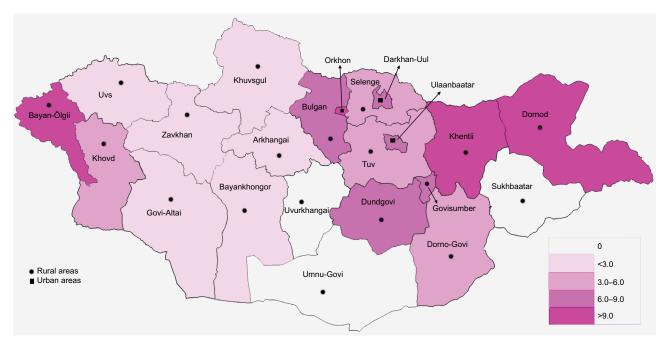


Figure 2 Age-standardized incidence rate by province, Mongolia, 2013.

Note: Data from Cancer Registry Data, National Cancer Center, Mongolia, 2009–2013.

available for collecting data across the whole of the country, data from rural areas may be impacted by limited health facilities, number of professionals and data distributions system compared with 3 urban locations in the country – the city of Ulaanbaatar (UB) and the larger provinces of Darkhan-Uul and Orkhon. A strong relationship between breast cancer and age has also been identified among Mongolian women with the highest incidence rate of 28.4/100,000 occurring among the 50–54 years old compared to 3.7 per 100,000 in the 15–39 age group.² Although the peak age appears to be closer to other Asian countries, at least a decade difference is noted when compared with westernized states (Figure 3).

Asia in general has a low incidence of breast cancer compared with westernized countries. However, compared with Mongolia the disease is still common among Asian women^{6,16–20} and with a mean incidence of 29 per 100,000 in 2012, it is the most common cancer.² Also when compared with countries such as the US, UK and Australia, the younger median age at diagnosis (40–50 years as opposed to 50–60 years) and the more progressed stage of the disease when diagnosed in Asia are quite different.^{17–19} However, due to lifestyle and reproductive changes in Asia, differences from westernized states to some extent are diminishing; for example, the incidence rates are increasing rapidly in the former, even though in absolute terms the rates for the time being remain low.^{20–22}

In the westernized world, breast cancer is the most common cancer among women with a mean incidence of 90 per 100,000 in 2012.² It has been observed that while age-standardized rates (ASRs) for breast cancer are higher, the rates are beginning to stabilize within developed countries unlike developing countries where breast cancer is continuing to emerge as an increasingly important health issue.^{23,24}

Breast cancer survival

In Mongolia, data on survival of breast cancer are very limited. With the lack of an early detection program and limited diagnostic and therapeutic resources, breast cancer is diagnosed at late stages, leading to a 5-year survival rate of 57% that is considerably lower than that described in other countries as highlighted by the recent global study based on the data between 2005 and 2009.25 Similar findings were shown in the report from the NCC for the period 2003-2013, which found that the average survival time for Mongolian patients with breast cancer was 3.2 years. 14 Also in Mongolia, 88.7% of women diagnosed with breast cancer are reported to die from the disease within 5 years, 14 which is significantly higher than westernized states where 5-year survival is closer to 90%. The limited access to health services in remote areas may play a role in delay of diagnosis resulting in poor survival outcome.

As a result of the advanced presentation of disease, the majority of breast cancer patients have radical mastectomy with axillary dissection combined with radiotherapy and chemotherapy, leading to a worse survival outcome compared

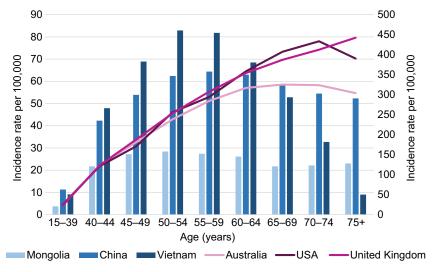


Figure 3 Age-specific incidence rate of female breast cancer, 2012.

Notes: The right axis indicates the incidence rates for Mongolia, China and Vietnam; the left axis indicates the incidence rates for Australia, USA and UK. Data from GLOBOCAN 2012.²

with the more developed treatments adopted elsewhere. Since early stage breast cancer is less frequent, the implementation of breast conserving surgery along with radiotherapy has only been conducted on a small proportion of patients. The use of radiation therapy remains low in Mongolia partially due to shortage of radiation oncologist and technologists. In addition, to treatment options, breast cancer survival can also be affected by other factors such as lifestyle, environmental, cultural and socioeconomic;²⁶ however, information on these factors and their association with survival is not well understood in Mongolia.

In Asia, inconsistency of early detection programs and the varying levels of access to appropriate treatment have led to the survival rates varying considerably across the region. In Japan, Republic of Korea and People's Republic of China, for example, 5-year survival rates of 85%, 83% and 82%, respectively, have been reported for invasive cancer cases during 2005-2009.25 More similar to Mongolia, survival rate is reported to be lower in Malaysia and India, where the rate is closer to 68% and 60%, respectively.²⁵ In addition, the majority of breast cancer patients from developing countries in Asia continue to be diagnosed at a relatively late stage and over 50% of all managed patients present with locally advanced cancer at the time of diagnosis. 21,27,28 It is important to note that even when detection and treatment opportunities exist, other factors, including estrogen receptor (ER) status, preexisting chronic disease and ethnicity impact upon variation in survival. For example, ER-negative cancers are more commonly reported among Asian women, which is associated with poor prognosis.19

The introduction of mammogaphic screening in westernized states highlights a potential way forward for Mongolia and other developing Asian states. Screening aimed at asymptomatic patients has led to marked improvements in overall survival. For example, In 1991, screening was introduced in Australia and 5-year survival rate was shown to increase from 72% to 89% between the periods 1982–1995 and 2006–2010.²⁹ However, the impact of changes in prevalence of risk factors and advances in diagnosis and treatment on favorable survival outcome cannot be ruled out. Such factors impact on stage of diagnosis and treatment. The former is a critically important determinant of survival, particularly regarding tumor size and the involvement of lymph nodes.^{30,31}

Breast cancer mortality

There is a paucity of quality data around mortality rates in Mongolia with some rural areas having no mortality data. A relatively recent national study¹³ demonstrated that 93.4% of the Mongolian women with breast cancer were diagnosed at advanced stages (Figure 4). The data that are available indicate that breast cancer is the 8th most common cause of cancer-related death in the country with age-standardized mortality rate of 2.7 per 100,000 in 2012. Comparisons with Asian and Westernized countries are demonstrated in Table 1. Similar to other developing Asian countries, mortality rates (and incidence numbers) are predicted to increase due to adoption of westernized lifestyle and recent increased detection and treatment facilities in Mongolia.

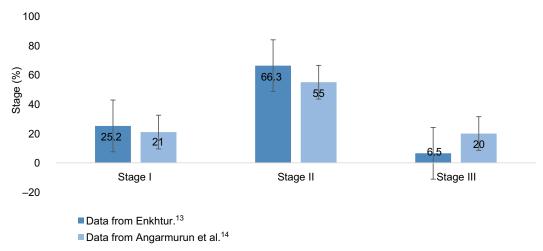


Figure 4 Breast cancer by stage, Mongolia.

Table I Estimated breast cancer incidence and mortality by country, 2012

Countries	Incidence		Mortality	
	Number	ASR (world)	Number	ASR (world)
Mongolia	125	9.4	50	1.6
Asia	650,983	29.1	231,013	6.4
Westernized countries*	127,016	89.5	26,906	7.3

Notes: The rates are age-standardized per 100,000 female population. *North America, Western and Northern Europe, Australia/New Zealand. Data from GLOBOCAN 2012.²

Abbreviation: ASR, age-standardized rate.

Although the ASRs for breast cancer in Mongolia historically have been consistent, a recent important increase has been described by recent national mortality data between 2009 and 2013 (Figure 5) where the rate was almost doubled from 1.4/100,000 in 2009 to 2.7/100,000 in 2012. Interestingly, more than half of the deaths from breast cancer were registered in the capital city of UB in 2013. This figure is likely to be explained at least in part by population distributions since almost 50% of the Mongolian people live in UB³² and also by an urban lifestyle, which reflects westernized activities. Following the 2009–2012 increase, the mortality rate was declined to 1.6/100,000 in 2013. The explanation for this unusual change to the trend is unclear and may reflect incomplete data coverage.

Regional mortality variations in Mongolia, however, are not confined to the capital. Outside UB, variation in mortality rates is also evident with numbers being higher in rural provinces such as Bayan-Ölgii (4.4) and Dornod (4.0), while being lower in other regions such as Khuvsgul (0.8) (Figure 6). While inconsistent registry data may have

contributed to these recorded variations, the stage of disease presentation is likely to be linked to these geographic variations.

While the rates are decreasing in many westernized nations, ^{29,33,34} mortality rates continue to increase in Asian countries, aligning with rapid increases in incidence of breast cancer. ^{19,22} In Asia, higher mortality rates are often observed in less developed countries despite these countries having low incidence rates. ^{18,28,35} A variety of explanations have been proposed for these regional-dependent patterns around culture, socioeconomic factor, geographical isolation, inadequate diagnostic equipment and poor treatment facilities; however, a consistent casual parameter is delayed presentation of the disease. ^{18,22,36}

In developed countries, breast cancer is the 2nd leading cause of cancer-related mortality after lung cancer, resulting in 198,000 deaths (15.4% of all cancer deaths) in 2012.² Mortality rates for breast cancer have steadily declined in most westernized countries;³⁷⁻⁴⁰ however, based on 2012 data, globally it is estimated that more than half million women will die from this disease annually, highlighting that breast cancer is not only the disease of developed countries.

Breast screening

Currently, in Mongolia, there is no population-based breast cancer screening; instead in 2010, guidelines were established for the prevention and management of the disease. As outlined in the guidelines, mammography is not recommended for early detection due to the relatively low incidence rates, limited resources for equipment and sparsely distributed populations in Mongolia.⁴¹ Therefore, clinical examination is the front line tool used for early

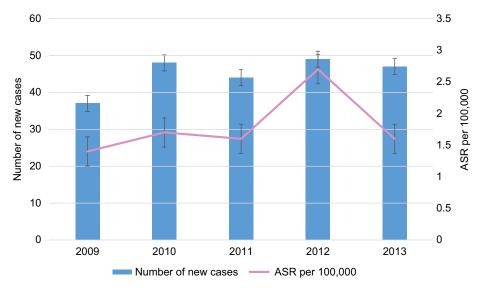


Figure 5 Mortality rate of female breast cancer, Mongolia, 2009–2013.

Note: Data from Cancer Registry Data, National Cancer Center, Mongolia, 2009–2013.

Abbreviation: ASR, age-standardized rate.

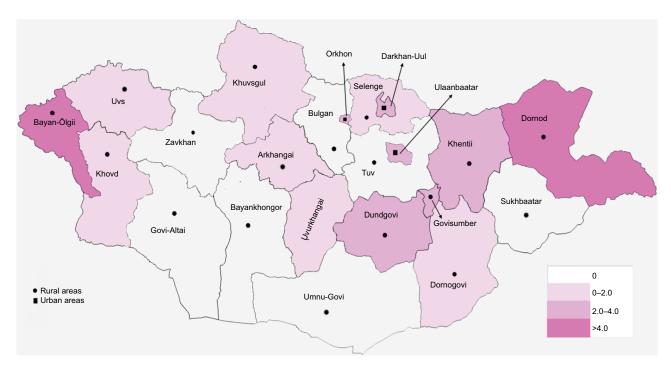


Figure 6 Age-standardized mortality rate by province, Mongolia, 2013. **Note:** Data from Cancer Registry Data, National Cancer Center, Mongolia, 2009–2013.

detection of the disease, and all suspected cases in local hospitals refer to the NCC for further diagnostic evaluation. However, this screening approach is having a limited impact, with only 1 in 3 women reporting breast self-assessment and only 3.2% and 1.7% undergoing clinical breast examination and mammography, respectively, as reported in a recent WHO review.⁴²

In recent years, the importance of early detection for breast cancer has become a serious issue in Mongolia due to increasing reports of late presentation and rapid increase in incidence of the disease. 12-14 Since mammographic facilities have been established in both public and private hospitals in Mongolia, implementation of breast screening could be an effective approach to detect the disease at early stages. Although mammographic screening reduces mortality in westernized countries, there is no evidence that systems used elsewhere would be effective in Mongolia. Therefore, more investigations around breast types and breast cancer

presentations in Mongolia need to be carried out so that a country-specific solution can be found.

In Asia, compared with the Western nations, breast cancer screening is not so widespread. The effectiveness of mammographic screening in Asia is the subject of debate due to the low incidence of breast cancer across the region, the prevalence of smaller breasts and denser breast parenchyma among Asian women. An Anonetheless, some Asian countries have adopted mammography for mass screening, such as Singapore, Japan and Taiwan, where some favorable outcomes have been reported. In Singapore for example (the first Asian country to introduce screening for 50–69 years old in 2002), the overall invasive cancer detection rate was reported to be 7.92 per 1000 women screened during 2002–2009 and a 12% increased rate of early stage cancers was recorded compared with prescreening period (before 2001).

In developed countries, breast screening with mammography is a well-established approach for the early detection of breast cancer and many experts agree that the benefit of mammography outweighs the harms. In particular, mammography is more accurate in older women than in young women. ^{49,50} The overall impact of screening is debated with reported reductions in mortality from 15% to 58%. ^{50–56}

Risk factor

The causes for breast cancer are not fully understood but women with certain risk factors are more likely to develop the disease than others. In Mongolia, the risk factors associated with breast cancer are understudied. The only available case-control study¹³ involved 522 breast cancer patients and was locally published in 2009. Similar to other countries, this limited evidence proposes the following risk factors: reproductive factors including early menarche, irregular menstrual cycles, nulliparity and obesity; induced abortion, which was aligned with a recent Chinese meta-analysis;⁵⁷ and various disease states including ovarian chronic inflammation and cysts and uterine fibroids. 13 It is also interesting to note that ethnic minority groups are expected to have low rates of incidence and mortality since these women generally adhere to more traditional lifestyles compared with their westernized counterparts; however, the opposite pattern was observed in Mongolia, which suggests a uniqueness around genetic and risk factors. While the aforementioned factors have been shown to be relevant to Mongolia, other important parameters reported for other countries such as family history, lifestyle factors and hormone replacement treatment (HRT) are currently poorly understood in Mongolia.

Current knowledge regarding breast presentation remains poor, which demands a better understanding of this disease among Mongolian women. For example, mammographic breast density has been shown to be one of the most important risk factors in westernized populations but nothing is known around this feature among Mongolian women. Westernized women with high density have been shown to have 4- to 6-fold increased lifetime risk of breast cancer.58-61 This is a feature that can be assessed from mammographic examinations and while mammography occurs in Mongolia, little data are gathered on density distributions, particularly across ethnicity, age, body mass index and menopausal status. The absence of information is not the case across Asia; for example in China, breasts are known to be fattier than in American women in older age but denser in young age. 62 The importance of understanding this feature for identifying risk and indeed for determining the optimal method of diagnostic imaging is emphasized so that prevention, early detection and intervention of the disease can be facilitated.

In Asia, while the prevalence and distribution of risk factors among Asian women vary between countries and studies, some common features have been noted in westernized studies. 19,63-66 A recent Asian meta-analysis summarized that more than 3 abortions, family history, late age at first live birth (over 30 years old), smoking, no breastfeeding, alcohol consumption and longer usage of HRT are associated with breast cancer among Asian women.⁶⁷ Other risk factors reported among these women include body fat, which reduces and increases the risk of breast cancer in premenopausal and postmenopausal women, respectively. 68,69 Uniquely to this region, however, a higher intake of soy food with their associated protective effect for breast cancer has been reported among Asian women, 70,71 and in China, consumption of green tea was reported to be associated with risk of breast cancer, 72 whereas this was not observed among Japanese women.73

In developed countries, certain lifestyle activities, westernized diets, high socioeconomic status, physical inactivity and obesity are reported to be associated with breast cancer risk in these regions.^{74–76} For example, a European cohort study revealed that an increased risk of 31% for breast cancer is strongly associated with obesity among postmenopausal women.⁷⁶ Family history and gene mutations such as BRCA1 and BRCA2 are also well known, established risk factors.^{77–81} In general, hormonal factors are different in prevalence between breast cancer subtypes; however,

hormone receptor-positive tumors were consistently being reported among Western women. 82,83 Westernized women also tend to have shorter breastfeeding period and low parity compared with Asian women, resulting in high risk for developing breast cancer. 84,85

In summary, what is distinct about Mongolia?

With the limited available data, this review is the first attempt to describe the current situation of breast cancer in Mongolia while investigating different aspects of the disease. It is evident that breast cancer incidence rate in Mongolia is among the lowest in the world and there are several possible explanations that have been considered in the above sections and relate to lifestyle, health policy and quality of registry data. With regard to lifestyle, reproductive activities such as longer breast feeding86 and higher number of full-term pregnancies probably play a significant role, and as described within the only peerreviewed publication¹² on the topic, dietary habits around high-level consumption of tea, meat and dairy product may also be key. However, absence of a breast screening program must also be a contributory factor particularly since it is evident that rates of breast cancer have increased in those Asian countries that have recently introduced formal screening programs. 48,87 Finally, reliability of cancer registry data cannot be excluded as incomplete data exist. Until screening procedures are formalized and accurate registry data become available, the importance of breast cancer to Mongolia cannot be fully appreciated. What is appreciated is that late disease presentation, poor survival and high mortality demand a coordinated effort between researchers, clinicians and health strategists to ensure that Mongolian women do not continue to suffer poorer health outcomes compared with their neighbors and westernized counterparts.

Conclusion

While breast cancer incidence is low in Mongolia compared with other Asian countries and westernized states, incidence rates are gradually increasing. Early detection with optimal imaging methods may be a key to minimizing the impact of this disease on Mongolian women; however, not enough is known about the type of cancer being presented, the regional distribution of the disease and the type of women most effected, which is needed to facilitate allocation of limited resources.

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Disclosure

The authors report no conflicts of interest in this work.

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