Breast cancer in Egypt, China and Chinese: statistics and beyond

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According to the latest report of The International Agency for Research on Cancer (GLOBOCAN 2012), breast cancer (BC) is by far the world's most common cancer among women, and the most likely cause that a woman will die from cancer worldwide (1).

The disease is reported as the most frequent cancer among women in 140 of 184 countries worldwide including China the world's most populous country (1,2). Notably, the incidence rates of BC varies dramatically across the globe, being always highest in more developed regions, namely North America, Western Europe (more than 90 new cases/10⁵ women annually), compared with less than 30/10⁵ women annually in regions like Eastern Asia (1). In the current issue of 7TD, Zeng et al. (3), have published the first national dataset on the incidence and mortality of female BC in China, for the year 2010, highlighting also some trends in incidence, and mortality, in different geographical regions within China. The authors should be commended for being the first to provide the literature with a wealth of data regarding BC statistics in China, which harbors around one fifth of all women around the world. According to the authors' estimations, the age standardized incidence rate (ASIR) of BC is 24.2/10⁵ and age standardized mortality rate (ASMR) is $6.36/10^5$. As such, these figures would confirm the known fact, that the mainland Chinese women have one of lowest BC incidence and mortality rates, when compared to the rest of the world.

Two important questions should be raised with this kind of findings: how far the authors could accurately quantify BC burden (occurrence and outcome) in China? and if their figures are true, then why the mainland Chinese women are so much protected against BC versus Chinese women in other Asian or western counties?

To start with, we need to emphasize that Zeng *et al.* analysis included only 12.96% of the female population of China. This means that the vast majority of Chinese women are not truly represented in their analysis, especially in the western region (less than 10% of their sample size), where the incidence is ~30% less than that reported in other areas. Obviously, "a cautious interpretation" is warranted with this kind of national epidemiological studies, when so many regions in a given country have low or no accurate registry data. Nonetheless, this is a common situation in many countries with emerging population-based registries (4). For instance, in Egypt; with its much smaller scale in geographical and population size, a network of five regional population-based registries, spreading over the whole country, was developed during the last two decades. For the sake of national cancer statistics, Egypt was divided into three regions namely upper, middle and lower Egypt. One or more defined states (governorates), with a regional population-based registry, were selected to represent each region (4). The data evolved on cancer incidence from each cancer registry were extrapolated in a similar methodology like that adopted by Chinese group. BC is estimated to be the most common female cancer in both China and Egypt, despite a marked difference in incidence rates, being much higher among Egyptian women compared to Chinese (ASIR are $48.8/10^5$ and $24.2/10^5$ respectively) (3,4). In the two countries BC incidence differs considerably among rural and urban populations, with the higher urban incidence being consistent across all women above the age of 45 years (3,5). A similar observation was also found in many developing countries like India, where BC incidence in rural registry of Barshi was 7.2/10⁵ compared to 31.3/10⁵ in adjoining city of Mumbai (6). In fact, the urban population in developing countries, might suffer from a higher exposure to xenoestrogens, which have been linked to evolution of hormone receptor positive BC in industrialized countries (7).

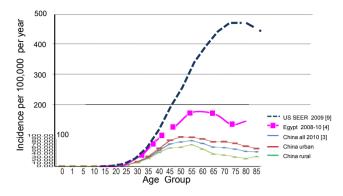


Figure 1 Age specific incidence rates of breast cancer in China, Egypt, and USA.

Pattern of age specific incidence of BC from east to west

Like other countries the age-specific incidence rates in both Egypt and China show a progressive increase after the age of 30 years, to reach a sharp peak of 80/10⁵ at the age group of 55-59 years in China, versus a sharp peak of 177.4/10⁵ at the age group of 60-64 years in Egypt (3,4). Importantly, in both countries there was a gradually decline in the age-specific incidence thereafter. This pattern of the age specific incidence is in clear contradistinction with western countries, where a continuous rise in BC incidence is observed with increasing age, to reach a peak of $\sim 450/10^5$, at the age 80 years, thus rendering the differences in age-specific incidence especially large in elderly women (8-10) (Figure 1).

We do not know whether this difference is completely real, or it is partially induced by under-reporting of BC cases specifically among elderly patients, in countries with emerging cancer registry programs. Notably, a pattern of BC age specific incidence, similar to that seen in China and Egypt, has been reported in some Asian countries with a completed national cancer registries like Japan and Singapore (11,12). This would certainly challenge the rule of "The Older the Age; The Higher the BC Risk", which might not be a default hypothesis, when it comes to North African and East Asian women.

Are the mainland Chinese women more protected from BC?

The reasons for the wide international variation in incidence rates of BC are not fully explained. However, it seems true that the low-risk Chinese (and other Asian) populations

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are enjoying a multitude of biological and lifestyle features that collectively provide a significant protection against the development of BC. In a cross sectional study, Chinese women were found to be significantly less obese (~14 kg less), have their menarche a vear later, are more often parous, and consume a lower western diet, compared to Caucasian women (13). The protective effect of Chinese diet was highly emphasized during the last decade. In one study on 1,602 BC women and 1,500 healthy women, all living in Shanghai, it was found that a "meat and sweet" western-style diet was independently associated with a 60% increased risk of BC, compared to a "soy and vegetable" traditional Chinese diet (14). The effect was more significant for estrogen-receptor positive tumors, where the risk of getting this phenotype of BC was almost doubled, among women utilizing western diet (14). A recent metaanalysis, found that the Chinese diet intake is associated with a 41% reduction in BC risk among Asian women (15). This main part explain why Chinese women migrating to North America, adopting a westernized lifestyle, will show a significant rise of BC incidence over a few generations (10,16). Taken together, lifestyle and socio-demographic factors stand as major players in the low BC burden, reported in China and other developing East Asian countries. However and in spite of the major similarities in lifestyle, the incidence rate of BC among Chinese women in the mainland is strikingly lower than that reported for Chinese women in other Asian countries. For example, in Singapore, where a complete national registry coverage is accomplished, an ASIR of $63.7/10^5$ has been reported among the Chinese population there (12). Similar figures were also reported in Hong Kong $(61/10^5)$ (17) and Malaysian Chinese (59.9/10⁵) (18), with a somewhat lower incidence among Taiwanese women $(44.45/10^5)$ (19) (*Table 1*).

It does not seem that, factors like life expectancy, more urbanization and presence of BC screening programs, can fully explain a 2-3 fold decrease in BC incidence among the mainland Chinese women, compared to Chinese women in other far east countries.

In conclusion, many factors can be directly responsible for the large variation in BC epidemiology across the globe; including life style, life expectancy, wide spread use of screening mammography and accuracy of national cancer statistics. The later factor is specifically crucial in developing countries, where the emerging population-based cancer registries may suffer from under-recording of some of the occurring BC (as well as other cancers). We believe that a wider expansion of the national cancer registries in both

 Table 1 Breast cancer ASIR and ASMR for East and West

 Chinese women

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Chinese population	ASIR⁺	ASMR ⁺⁺
Chinese Mainland [3]	24.2	6.36
Chinese Singapore [12]	63.7	14.2
Chinese Hong Kong [17]	59.9	NA*
Chinese Malaysia [18]	61	9.1
Chinese Taiwan [19]	44.45	NA*
Chinese US [16]	73.5	11.5 [10]**

⁺ASIR, age standardized incidence rate; ⁺⁺ASMR, age standardized mortality rate; ^{*}NA, not available in the published reference; ^{**}, for all Asian-Americans.

China as well as Egypt, will be able to answer some of the many questions, related to variations in BC statistics in the developing countries.

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