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# Epidemiological trends of women's cancers from 1990 to 2019 at the global, regional, and national levels: a population-based study



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### **Abstract**

**Background:** Every year around the world, more than 2 million women are diagnosed with breast cancer and genital tract cancers. However, there are rare studies comprehensively describing the global and regional trends of incidence and mortality of women's cancers.

**Methods:** To study the burden and trend of women's cancers, we conducted this cross-sectional study based on the epidemiologic data of Global Burden of Disease 2019. In this study, female patients with breast cancer, cervical cancer, ovarian cancer, and uterine cancer worldwide from 1990 to 2019 were involved. The incidence, death, and disability-adjusted life-year (DALY) were used to measure the outcomes of women's cancers. The estimated annual percentage change (EAPC) was calculated to assess the changing trend of cancer burden.

**Results:** Among the four women's cancers, the burden of female breast cancer was highest. During the past 30 years, the incidence, death, and DALY of female breast cancer kept increasing worldwide. In most regions especially developing countries, cervical cancer was the second most common women's cancer. At the same time, ovarian cancer and uterine cancer occurred less frequently. Generally, the age-standardized incidence rates (ASIRs) of breast cancer, ovarian cancer, and uterine cancer were positively correlated to sociodemographic index (SDI) value. In contrast, the ASIR of cervical cancer was negatively correlated to SDI value.

**Conclusions:** Our study indicates that the incidence and mortality of women's cancers have geographical variations and change along with SDI value. The results might be helpful to policy-makers to allocate healthy resources to control women's cancers.

**Keywords:** Breast cancer, Cervical cancer, Ovarian cancer, Uterine cancer, Gynecological cancer, Cancer epidemiology

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### **Background**

According to GLOBOCAN 2020 Project data, breast cancer, cervical cancer, ovarian cancer, and uterine cancer are among the top 10 common cancers in females worldwide [1]. It is estimated that female breast cancer has been the most common cancer, surpassing lung cancer globally [1]. There are significant geographical variations in the incidence rate of female breast cancer [1]. The incidence rate of female breast cancer in some developed regions (Australia, New Zealand, North America) is nearly three times higher than that in developing regions (South-central Asia, Middle Africa, East Africa) [1]. Also, among females, breast cancer is the leading cause of cancer deaths in most countries [1]. However, in some sub-Saharan Africa countries, breast cancer is preceded by cervical cancer, which causes the most cancer deaths [1]. Although cervical cancer could be prevented by HPV vaccination and screening pre-cancerous lesions, cervical cancer's incidence and mortality are still high in these counties due to lacking essential health interventions [2]. Relative to cervical cancer, ovarian cancer and uterine cancer occur less frequently globally [3-5]. Generally, the incidences and deaths of four types of women's cancers have continued to grow during the past decades.

Global Burden of Disease 2019 (GBD 2019) study provides the epidemiological data of 369 diseases and injuries in 204 countries or territories from 1990 to 2019, which updates based on GBD 2017 [6-8]. The incidences, deaths, disability-adjusted life-year (DALY) of diseases could be used to measure disease and economic burdens [9–14]. The epidemiological patterns of women's cancers vary in different regions and change over time. A systematic analysis comprehensively reporting the variation trends of women's cancers helps policymakers measure the burden of women's cancer, build health service infrastructures, and allocate public health resources. In this cross-sectional study, we reported the incidences, deaths, and disability-adjusted life-years (DALYs) of female breast cancer, cervical cancer, ovarian cancer, and uterine cancer from 1990 to 2019 in 204 countries. Besides, we analyzed the correlation between incidence rate or mortality rate and sociodemographic index (SDI).

### **Methods**

### **Study Population and Data Collection**

All data were obtained from GBD 2019 study, including incidences, deaths, DALY, age-standardized rate of incidence (ASIR), age-standardized rate of death (ASDR), and age-standardized rate of DALY from 1990 to 2019. SDI value is a parameter to evaluate a country's development status, which is scored from 0 to 1 by calculating the geometric mean of fertility rate, length of education,

and per capita income [15]. All countries or territories are classified into five groups according to SDI value: low SDI, low-middle SDI, middle SDI, high-middle SDI, and high SDI. SDI values of all countries from 1990 to 2019 were obtained from Institute for Health Metrics and Evaluation.

Because the study was based on publicly available dataset, this study was exempted by the ethics committee of Tongji Hospital, Tongji Medical College, Huazhong University of Science and Technology. This study followed the guideline of cross-sectional study described in the Guidelines for Accurate and Transparent Health Estimates Reporting (GATHER) [16].

### Statistical Analysis

Firstly, we analyzed the burden of women's cancers at the global, geographic regional, and national levels. Additionally, to offset the differences in demographic and population number, we used the age-standardized rates of incidence, death, and DALY to explore incidence and mortality levels in different regions and countries. All rates were presented as numbers of per 100,000 persons.

Moreover, we assessed the variation trends of incidence and mortality by calculating estimated annual percentage changes (EAPCs) of age-standardized rates. In the calculating formula  $y = \beta x + \alpha$ , x refers to calendar year, and y refers to log10 (age-standardized rate). EAPC was calculating as following: EAPC =  $100^*$  ( $10^5$  -1) [15, 17-19]. When EAPC value and its 95% confidence interval (CI) are greater than zero, the age-standardized rate is upward. Conversely, the age-standardized rate is downward [20].

Besides, to investigate the potential effect of SDI on cancer burden, we analyzed the Spearman correlation between SDI value and age-standardized rates of incidence, death, and DALY [21]. All statistical analyses were conducted using R software (version 4.0.0, R Project for Statistical Computing). A two-tailed *P* value below 0.05 was regarded as statistical significance. Data visualization was performed using R software with package ggplot2.

### Results

### Female breast cancer

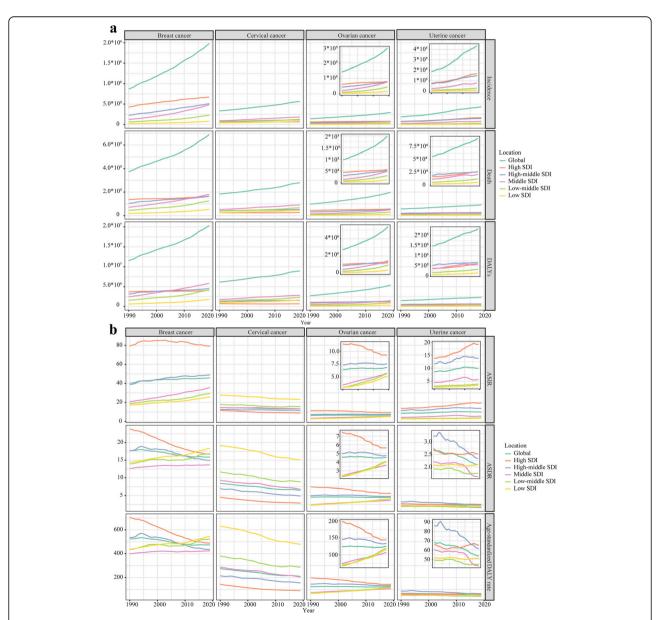
During the past 30 years worldwide, the incidences of female breast cancer increased from  $867.62 \times 10^3$  (95 %CI  $840.4 \times 10^3 \sim 894.76 \times 10^3$ ) to  $1977.21 \times 10^3$  (95 %CI  $1807.61 \times 10^3 \sim 2145.21 \times 10^3$ ) cases (Table S1). At the same time, the deaths of female breast cancer increased from  $375.02 \times 10^3$  (95 %CI  $358.98 \times 10^3 \sim 390.82 \times 10^3$ ) to  $688.56 \times 10^3$  (95 %CI  $635.32 \times 10^3 \sim 739.57 \times 10^3$ ) cases (Table S2), and the DALYs of female breast cancer increased from  $11526.68 \times 10^3$  ( $11021.13 \times 10^3 \sim 12107.83 \times 1$ 

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 $10^3)$  to  $20310.19\times10^3$  (95 %CI 18744.8  $\times$   $10^3\sim21866.65\times10^3)$  years (Table S3). Globally, from 1990 to 2019, the incidences, deaths, and DALYs of female breast cancer were gradually increased (Fig. 1a). For age-standardized rates, the ASIR of female breast cancer rose from 40.12 (95 %CI 38.78  $\sim$  41.33) in 1990 to 45.86 (95 %CI 41.91  $\sim$  49.76) in 2019, with EAPC<sub>ASIR</sub>: 0.36 (95 %CI 0.31  $\sim$  0.42) (Fig. 1b). The ASDR of female breast cancer decreased from 17.76 (95 %CI 16.93  $\sim$  18.51) in 1990 to 15.88 (95 % CI 14.66  $\sim$  17.07) in 2019, with EAPC<sub>ASDR</sub>: -0.51 (95 %CI -0.57 $\sim$ 0.46). The age-standardized DALY rate of female breast cancer decreased from 524.87 (95 %CI 501.78  $\sim$  551.15) in

1990 to 473.83 (95 %CI 437.3  $\sim$  510.51) in 2019, with EAPC<sub>age-standardized DALY rate</sub>: -0.51 (95 %CI -0.57 $\sim$ -0.45).

The burden of female breast cancer varied in different regions. In 2019, the incidence number of female breast cancer was highest in high SDI region (673.15  $\times$  10³, 95 %CI 601.27  $\times$  10³  $\sim$  747.67  $\times$  10³). However, the ASIR of female breast cancer was stable in high SDI region (79.3 with 95 %CI 77  $\sim$  80.87 in 1990; 79.22 with 95 %CI 70.83  $\sim$  87.7 in 2019; EAPC<sub>ASIR</sub>: -0.12, 95 %CI -0.22  $\sim$  -0.01). In contrast, the ASIR of female breast cancer was rapidly increased in middle SDI region (20.81 with 95 %CI 19.25  $\sim$  22.45 in



**Fig. 1** The changing trend of the burden of women's cancers from 1990 to 2019. **a** The changing trends of incidence, death, and DALYs. **b** The changing trends of ASIR, ASDR, and age-standardized DALY rate. Note: DALY, disability adjusted life year; ASIR, Age-standardized incidence rate; ASDR, Age-standardized death rate

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1990; 35.52 with 95 %CI 31.47  $\sim$  39.81 in 2019; EAP-C<sub>ASIR</sub>: 1.87, 95 %CI 1.84  $\sim$  1.9). Relative to ASIR, at the same time, the ASDR and age-standardized DALY rate were significantly declined in high SDI region: EAPC<sub>ASDR</sub>: -1.36, 95 %CI -1.41 $\sim$ -1.31; EAPC<sub>age-standardized DALY rate</sub>: -1.41, 95 %CI -1.47 $\sim$ -1.36.

In 1990, the USA, China, Germany were the top three countries with the most incidence cases. In 2019, China, the USA, and India were the countries with the most incidence cases  $(368.37 \times 10^3, 251.53 \times 10^3, 144.09 \times 10^3;$  respectively) (Fig. 2a). In 2019, China, India, and the USA were the countries with the most deaths  $(93.50 \times 10^3, 82.10 \times 10^3, 54.40 \times 10^3;$  respectively) (Fig. 2b) and most DALYs  $(2877.24 \times 10^3, 2659.24 \times 10^3, 1387.67 \times 10^3;$  respectively) (Fig. 2c). Monaco was the country with the highest ASIR in 2019 (149.60, 95) %CI  $108.67 \sim 193.81$ ). The ASIR

was relatively high in the USA in 2019 (94.21, 95 %CI 77.35 ~ 115.12) (Figure S1a). Solomon Islands was the country with the highest ASDR and age-standardized DALY rate in 2019 (75.04, 95 %CI  $59.00 \sim 94.48$ ; 2635.74,  $2026.64 \sim 3377.14$ ; respectively). Pakistan had a relatively high ASDR and age-standardized DALY rate in 2019 (51.94, 95 %CI  $39.03 \sim 69.76$ ; 1570.06,  $1177.2 \sim 2135.47$ ; respectively) (Figure S1b and Figure S2c).

### Cervical cancer

Globally, the incidence cases of cervical cancer increased from  $335.64 \times 10^3$  (95 %CI  $300.35 \times 10^3 \sim 393.89 \times 10^3$ ) in 1990 to  $565.54 \times 10^3$  (95 %CI  $481.52 \times 10^3 \sim 636.43 \times 10^3$ ) (Table S4). In the same period, the deaths of cervical cancer increased from  $184.53 \times 10^3$  (95 %CI  $164.84 \times 10^3 \sim 218.94 \times 10^3$ ) to

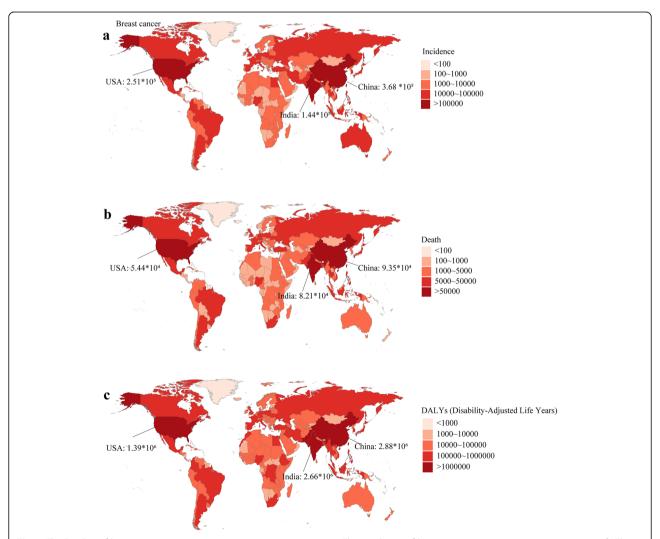


Fig. 2 The burden of breast cancer in 204 countries or territories in 2019. a The incidence of breast cancer in 204 countries or territories; b The death of breast cancer in 204 countries or territories; c The DALYs of breast cancer in 204 countries or territories. Note: DALY, disability adjusted life year

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 $280.48 \times 10^3$ (95 %CI  $238.86 \times 10^3 \sim 313.93 \times 10^3$ ) (Table S5), and the DALYs of cervical cancer increased from  $6176.25 \times 10^3$ (95 %CI 5437.67 ×  $10^3 \sim 7316.93 \times 10^3$ to  $8955.01 \times 10^3$ (95 %CI  $7547.73 \times 10^{3} \sim 9978.46 \times 10^{3}$ ) (Table S6). The incidences, deaths, and DALYs of cervical cancer kept steadily increased from 1990 to 2019 in the world (Fig. 1a). However, the global incidence rate and mortality rate gradually decreased in the past three decades: EAPC<sub>ASIR</sub>: -0.38, 95 %CI -0.41~-0.34; EAPC<sub>ASDR</sub>: -0.93, 95 %CI -0.98~-0.88; EAPC<sub>age-stan-</sub> dardized DALY rate: -0.95, 95 %CI -1.00~-0.89 (Fig. 1b).

Middle SDI region had the highest cervical cancer burden in 2019:  $183.34 \times 10^3$  (95 %CI  $144.49 \times$  $10^3 \sim 208.86 \times 10^3$ ) incidence cases,  $90.1 \times 10^3$  (95 %CI  $71.33 \times 10^3 \sim 103.2 \times 10^3$ ) deaths, and  $2817.25 \times 10^3$  $(2223.19 \times 10^3 \sim 3217.72 \times 10^3)$  DALYs. Low SDI region had the highest incidence rate and mortality rate in 2019: ASIR: 23.21, 95 %CI 18.31 ~ 28.76; ASDR: 15.05, 95 %CI 11.92 ~ 18.46; age-standardized DALY rate: 477.53, 95 %CI 374.33 ~ 591.38. High SDI region had the lowest incidence rate and mortality rate in 2019: ASIR: 8.91, 95 %CI 7.74 ~ 9.99; ASDR: 2.9, 95 %CI 2.6 ~ 3.1; age-standardized DALY rate: 89.72, 95 %CI 81.88 ~ 95.85. Besides, high SDI region had the most rapid decline in incidence rate and mortality rate: EAPCASIR: -0.95, 95 %CI -1.06~-0.85; EAPC<sub>ASDR</sub>: -1.57, 95 %CI -1.68~-1.46; EAPC<sub>age-standardized DALY rate</sub>: -1.62, 95 %CI  $-1.75 \sim -1.49$ .

China, India, Brazil were top three countries with the most incidence cases in 2019 (109.76  $\times$  10<sup>3</sup>,  $84.98 \times 10^3$ ,  $22.65 \times 10^3$ , respectively) (Fig. followed by the USA  $(19.11 \times 10^3$ , ranking fourth). Similarly, China, India, Brazil were top three countries with the most deaths and DALYs in 2019 (deaths:  $53.44 \times 10^3$ ,  $45.45 \times 10^3$ ,  $11.07 \times 10^3$ , respectively; DALYs:  $1622.24 \times 10^3$ ,  $1554.49 \times 10^3$ ,  $348.42 \times 10^3$ , respectively), followed by the USA (deaths:  $7.99 \times 10^3$ , ranking fifth; DALYs:  $224.78 \times 10^3$ , ranking sixth) (Fig. 3b and c). Kiribati had the highest agestandardized rates of incidence, death, and DALY in 2019 (ASIR: 108.8, 95 %CI 78.79 ~ 140.73; ASDR: 69.52, 95 %CI 51.09 ~ 88.92; age-standardized DALY rate: 2143.06, 95 %CI 1534.43 ~ 2781.26). In general, the incidence rate and mortality rate were low in China (ASIR: 11.01, ASDR: 5.13, age-standardized DALY rate: 157.5), India (ASIR: 13.1, ASDR: 7.38, age-standardized DALY rate: 239.49), and the USA (ASIR: 8.67, ASDR: 3.05, age-standardized DALY rate: 98.77) (Figure S2).

### Ovarian cancer

Relative to incidence cases of female breast cancer and cervical cancer, the incidence cases of ovarian cancer were much lower worldwide. In 2019, the incidence of ovarian cancer was  $294.42 \times 10^3$  (95 %CI  $260.65 \times 10^3 \sim 329.73 \times 10^3$ ) worldwide (Table S7). The deaths of ovarian cancer were  $198.41 \times 10^3$  (95 %CI  $175.36 \times 10^3 \sim 217.66 \times 10^3$ ), and the DALYs of ovarian cancer were  $5359.74 \times 10^3$  ( $4692.95 \times 10^3 \sim 5954.99 \times 10^3$ ) in 2019 globally (Table S8 and Table S9). From 1990 to 2019, the incidence and mortality rates of ovarian cancer remained stable at the global level (EAPC<sub>ASIR</sub>: 0.11, EAPC<sub>ASDR</sub>: -0.11, EAPC<sub>age-standardized DALY rate</sub>: -0.08).

Although high SDI region had the highest ovarian cancer burden and mortality from 1990 to 2019, the agestandardized rates of incidence, death, and DALY in high SDI region were promptly decreased: EAPC<sub>ASIR</sub>: -0.87, 95 %CI -0.96~-0.78; EAPC<sub>ASDR</sub>: -1.1, 95 %CI -1.17~-1.02; EAPC<sub>age-standardized DALY rate</sub>: -1.27, 95 %CI -1.34~-1.2. In contrast, despite having a relatively low ovarian cancer burden and mortality, low-middle SDI region and low SDI region had the most rapid increase in age-standardized rates of incidence, death, and DALY (Table S8 and Table S9).

China, India, and the USA were the top three countries with the most ovarian cancer incidence cases (China:  $45.48 \times 10^3$ , India:  $31.02 \times 10^3$ , the USA:  $26.79 \times 10^3$ ) (Fig. 4a), deaths (China:  $29.09 \times 10^3$ , India:  $22.35 \times 10^3$ , the USA:  $19.5 \times 10^3$ ) (Fig. 4b), and DALYs (China:  $835.06 \times 10^3$ , India:  $657.74 \times 10^3$ , the USA:  $426.5 \times 10^3$ ) in 2019 (Fig. 4c). Monaco had the highest ASIR (22.75, 95 %CI  $16.73 \sim 28.89$ ) and ASDR (13.67, 95 %CI  $9.78 \sim 17.24$ ), and Pakistan had the highest age-standardized DALY rate (348.37, 95 %CI  $182.72 \sim 562.44$ ) in 2019. Among China, India, and the USA, the USA had a relatively high age-standardized rates of incidence, death, and DALY in 2019 (Figure S3).

### **Uterine** cancer

Around the world, the incidence cases of uterine cancer elevated from  $187.19\times10^3~(174.63\times10^3\sim196.03\times10^3)$  in 1990 to  $435.04\times10^3~(397.02\times10^3\sim479.73\times10^3)$  in 2019 (Table S10). In the same period, the global deaths increased from  $56.13\times10^3~(95~\text{WCI}~51.1\times10^3\sim60.2\times10^3)$  to  $91.64\times10^3~(95~\text{WCI}~82.39\times10^3\sim101.5\times10^3)$ , and the global DALYs increased from  $1483.29\times10^3~(95~\text{WCI}~1317.51\times10^3\sim1612.75\times10^3)$  to  $2329.07\times10^3~(95~\text{WCI}~2092.95\times10^3\sim2560.89\times10^3)$  (Table S11 and Table S12). At the global level, the incidence rate of uterine cancer was increased, but the mortality rate was decreased during the past three decades (EAPC\_ASIR: 0.69, 95 %CI 0.57 $\sim0.81$ ; EAPC\_ASDR: -0.85, 95 %CI -0.93 $\sim$ -0.76; EAPC\_age\_standardized\_DALY rate: -0.84, 95 %CI -0.93 $\sim$ -0.75).

High SDI region had the most uterine incidence cases  $(168.02 \times 10^3, 95 \text{ }\%\text{CI } 148.61 \times 10^3 \sim 188.55 \times 10^3)$  in 2019 and the most rapidly increased ASIR (EAPC<sub>ASIR</sub>: 1.37,

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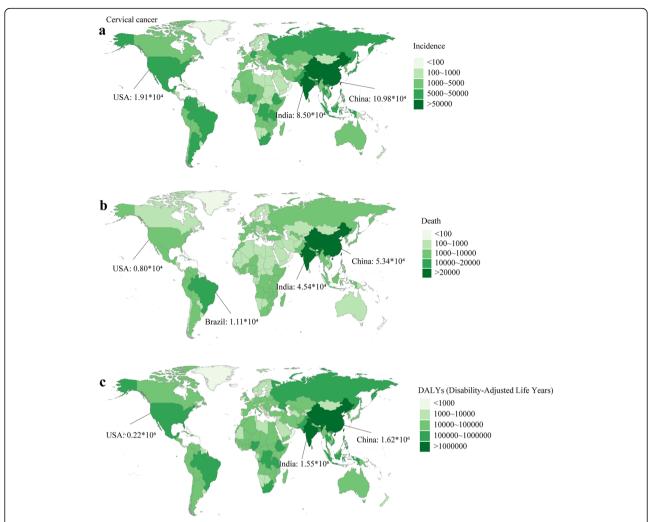


Fig. 3 The burden of cervical cancer in 204 countries or territories in 2019. **a** The incidence of breast cancer in 204 countries or territories; **b** The death of cervical cancer in 204 countries or territories; **c** The DALYs of cervical cancer in 204 countries or territories. Note: DALY, disability adjusted life year

95 %CI  $1.28 \sim 1.46$ ) from 1990 to 2019. Also, high SDI region had the most deaths  $(26.63 \times 10^3, 95 \text{ %CI } 24 \times 10^3 \sim 28.14 \times 10^3)$  and the highest ASDR  $(2.52, 95 \text{ %CI } 2.32 \sim 2.64)$  and age-standardized DALY rate  $(65.31, 95 \text{ %CI } 60.99 \sim 69.65)$  in 2019.

The USA, China, and Russia were the top three countries with the most incidence cases  $(80.07\times10^3,\,66.74\times10^3,\,42.71\times10^3,\,$  respectively) in 2019, followed by India  $(15.26\times10^3,\,$  ranking sixth) (Fig. 5a). China, the USA, India had the most deaths  $(12.22\times10^3,\,$   $10.26\times10^3,\,$   $7.04\times10^3,\,$  respectively) and DALYs  $(364.28\times10^3,\,$   $249.56\times10^3,\,$   $180.91\times10^3)$  in 2019 (Fig. 5b and Fig. 5c). Northern Mariana Islands had the highest ASIR  $(32.77,\,$  95 %CI  $21.3\sim42.36)$ , Grenada had the highest ASDR  $(11.3,\,$  95 %CI  $9.79\sim12.99)$ , and American Samoa had the highest age-standardized DALY rate  $(285.41,\,$  95 %CI  $185.19\sim370.95)$  in 2019. Compared to China and India, the USA had a higher age-standardized rates of

incidence, death, and DALY in 2019 (Figure S4). The age-standardized rates of women's cancers in all countries or territories were summarized on Table S13.

# The incidence, death, and DALY of women's cancers and demographic

In 2019, there were rare women's cancer patients aged less than 25 worldwide (Figure S5a). For breast cancer, the global incidence case and death number reached the peak at the age of  $50 \sim 70$ . Besides, the death number of breast cancer reached the second peak at the age older than 80, and the DALY of breast cancer reached the peak at the age of  $50 \sim 55$ . The ASIR and ASDR of breast cancer increased with age, reaching the peak at the age older than 80 (Figure S5b). The age-standardized DALY rate elevated with age as well, but it reached the plateau after the age of 55.

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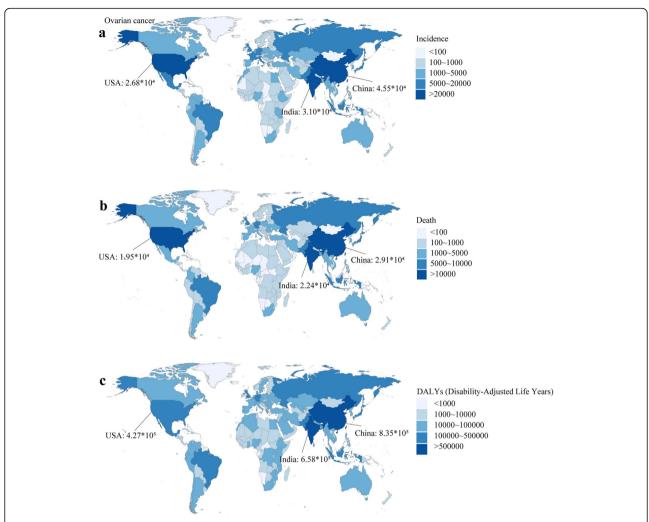


Fig. 4 The burden of ovarian cancer in 204 countries or territories in 2019. a The incidence of ovarian cancer in 204 countries or territories; b The death of ovarian cancer in 204 countries or territories; c The DALYs of ovarian cancer in 204 countries or territories. Note: DALY, disability adjusted life year

The global incidence of cervical cancer was in peak at the age of  $40 \sim 55$  (Figure S5a). The global death of cervical cancer reached a peak at the age of  $50 \sim 60$ . The global DALY of cervical cancer peaked at the age of  $50 \sim 55$ . The global ASIR of cervical cancer reached the plateau after the age of 40, and the global ASDR of cervical cancer increased with age. The global age-standardized DALY rate peaked at the age of  $55 \sim 60$  (Figure S5b).

The global incidence of ovarian cancer reached a plateau after the age of 50 (Figure S5a). The global death of cervical cancer climbed the first peak at the age of  $65 \sim 70$  and reached the second peak after 80. The global DALY of ovarian cancer was in peak at the age of  $50 \sim 65$ . The global ASIR and ASDR of ovarian cancer increased with age. The global age-standardized DALY rate of ovarian cancer peaked at the age of  $65 \sim 75$  (Figure S5b).

The global incidence of uterine cancer was in peak at the age of  $55 \sim 65$  (Figure S5a). The global death of uterine

cancer reached the first peak at the age of  $65 \sim 70$  and reached the second peak after the age of 80. The global DALY of uterine cancer peaked at the age of  $60 \sim 65$ . The global ASIR and age-standardized DALY rate of uterine cancer were in peak at the age of  $70 \sim 75$  (Figure S5b). The global ASDR of uterine cancer increased with age.

Generally, the demographic characteristics of women's cancer patients were similar in regions with different SDI values except for the ratio of patients aged older than 80 (Figure S6, S7, S8, S9, S10, S11, S12, S13, S14, S15). In high SDI region, there was a much higher ratio of patients aged older than 80, especially compared to the data of low SDI region.

# The correlations between SDI and rates of incidence and mortality

The ASIRs of breast cancer were significantly positively correlated with SDI value ( $\rho$ : 0.79, P < 0.0001) (Fig. 6). The

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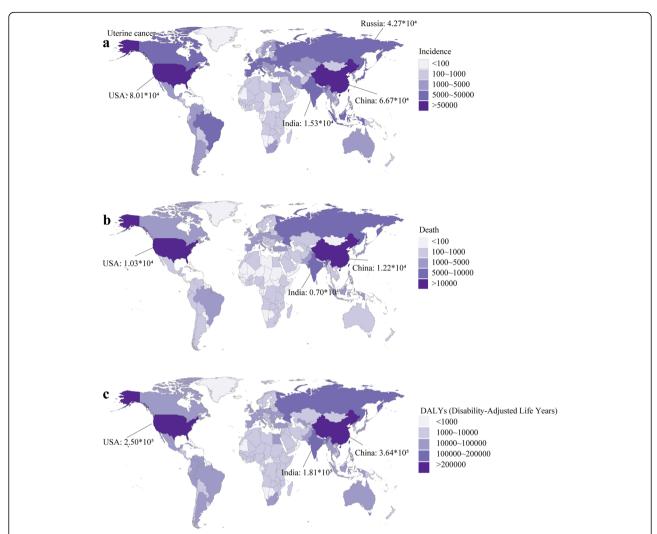


Fig. 5 The burden of uterine cancer in 204 countries or territories in 2019. a The incidence of uterine cancer in 204 countries or territories; b The death of uterine cancer in 204 countries or territories; c The DALYs of uterine cancer in 204 countries or territories. Note: DALY, disability adjusted life year

ASDR and age-standardized DALY rate of breast cancer exhibited non-linear correlations with SDI value. When SDI value was greater than 0.7, the ASDR and age-standardized DALY rate of breast cancer sharply decreased with the increase of SDI. In contrast, for cervical cancer, the age-standardized rates of incidence, death, and DALY were negatively correlative with SDI value ( $\rho$ : -0.61, P < 0.0001;  $\rho$ : -0.76, P < 0.0001; respectively).

For ovarian cancer, generally, the age-standardized rates of incidence, death, and DALY were positively correlative with SDI value ( $\rho$ : 0.86, P < 0.0001;  $\rho$ : 0.74, P < 0.0001;  $\rho$ : 0.73, P < 0.0001; respectively). However, when SDI was more than 0.7, the ASIR, ASDR and age-standardized DALY rate rapidly declined with SDI increase. For uterine cancer, the ASIR was significantly positively correlated with SDI value ( $\rho$ : 0.79, P < 0.0001).

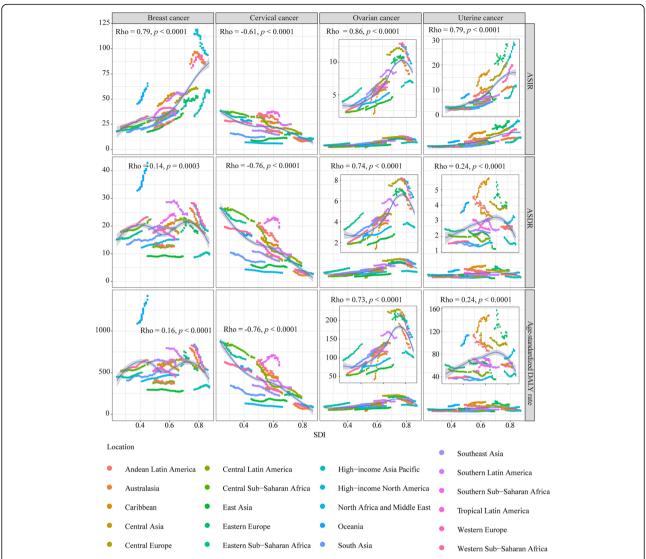
The correlation between mortality rate and SDI in uterine cancer was similar to that in ovarian cancer.

### Discussion

In the present study, we reported the incidences and mortalities of female breast cancer, cervical cancer, ovarian cancer, and uterine cancer. Around the world, the total incidence cases and death cases of the four types of cancers increased sharply from 1990 to 2019. Excluding the increased population and changed demographic, the ASIRs and ASDRs of the four types of cancers exhibited different epidemiological patterns.

Globally, the ASIR of breast cancer gradually increased, while the ASDR of breast cancer steadily decreased in the past 30 years. The two contradictory trends are partly attributed to the widely-adopted

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**Fig. 6** The correlation analyses of age-standardized rates and SDI from 1990 to 2019. The correlations between ASIR/ASDR/age-standardized DALY rate and SDI were analyzed based the data of 21 regions. Note: ASIR, Age-standardized incidence rate; ASDR, Age-standardized death rate; DALY, disability adjusted life year

mammographic screening and improved treatment strategy [22]. A pervious study based on GBD 2017 reported a similar conclusion: steadily growing ASIR and declining ASDR [15]. From 2017 to 2019, China, the USA, and India were still the top 3 countries with the most breast cancer incidence and death. Moreover, Li et al. found that high body mass index, alcohol use, and high fasting plasma glucose were risk factors of breast cancer-caused mortality [15]. In parallel, a registry-based cohort study in Norway showed that alcohol use and tobacco smoking were associated with higher risk of breast cancer for women with benign breast diseases [23]. Up to now, it has been identified multiple modifiable risk factors of breast cancer including alcohol, obesity, younger age of

menarche, older age of menopause, fewer childbearing, fewer breastfeeding, lower physical activity, and unhealthy lifestyles such as high fat consumption and low intakes of fruits, vegetables, and grains [24]. The alterations in reproductive patterns (such as low fertility rate), lifestyles, obesity rate, and age of menarche and menopause contribute to the increased incidence rate of breast cancer globally [25–28]. Besides, the breast cancer incidence rate varies in different ethnicities. It was reported that the incidence rate of white women was higher than that of ethnic minorities [29].

In the study, for cervical cancer, the age-standardized rates of incidence, death, and DALY were negatively correlated with SDI. Relatively, the incidence rate and death Yi et al. Biomarker Research (2021) 9:55 Page 10 of 12

rate were significantly higher in these low-source countries. It has been confirmed that the carcinogenic HPV infection is the primary cause of cervical cancer [30]. In the past decades, due to the implementation of HPV vaccination and HPV-based screening, the incidence rate of cervical cancer kept declining [31]. Also, based on the microscopic inspection of cervical scraping smear, the treatment of pre-cancerous lesions is the secondary prevention of cervical cancer [31]. However, in some developing areas, such as Southern Sub-Saharan Africa, it is hard for most women to access effective interventions [32]. Therefore, it is urgent for these areas to implement essential prevention and population-based screening programs, including introducing vaccination, screening cervical cancer and precursor lesions. It is estimated that the implementation of WHO cervical cancer estimation strategy could significantly decrease cervical cancer mortality by 99 % in the following century [33].

Our results showed that the incidence rate of ovarian cancer varied geographically, positively related to SDI value. Fewer breastfeeding, infertility or nulliparity, hormone treatment, and obesity are risk factors of ovarian cancer [34–38]. Contrarily, oral contraceptive is a strong protective factor for ovarian cancer [39, 40]. Previous studies showed that oral contraceptives could inhibit carcinogenesis by interfering with estradiol production and decreasing estradiol exposure in the menstrual cycle [39]. The high ovarian cancer incidence rate in regions such as Western Europe and North America, might be related to the high prevalence of these risk factors. In some developed countries, including the USA, the declined incidence rate paralleled the decreased mortality rate in the past 30 years. The developments in ovarian cancer treatment, including targeted therapy, neoadjuchemotherapy, intraperitoneal chemotherapy, aggressive surgery, contribute to the declined mortality rate [41].

The incidence rate of uterine cancer was mushrooming globally, especially in high SDI region. According to GBD 2019, in the USA, the ASIR of uterine cancer increased from 19.63 to 1990 to 28.80 in 2019. Obesity was a vital risk factor contributing to the increased uterine cancer incidence [42]. Overweight or obese women are more likely to develop uterine cancer than normalweight women [43]. Besides, decreased physical activity and a higher prevalence of diabetes are also the risk factors of uterine cancer [44]. Public health programs to help women maintain normal weight and increase physical activity could decrease uterine cancer risk.

Although population screening might be an effective measure to control cancer-caused mortality, the effectiveness of general population screening for women's cancers needs further validation in the real world. A recent randomized controlled trial in UK showed that annual transvaginal ultrasound screening increased the incidence rate of Stage I/II ovarian and tubal cancer but decreased the incidence rate of Stage III/IV ovarian and tubal cancer, relative to non-screening [45]. Notably, there was no significant reduction in disease-specific death in annual transvaginal ultrasound screening group, indicating the general population screening for ovarian cancer might be unnecessary at present stage [45].

Cancer is an aging-related disease. In the four types of women's cancers, we found the ASIR was positively correlated with age. Due to the demographic alterations in the future including the declining parity and the aging of baby boom generation, the mean age of some countries will further grow. It is estimated that in the USA, the incidence cases of ovarian cancer will increase by 37 %, from 20,921 cases in 2010 to 28,591 cases in 2030 [46]. This rise in ovarian cancer cases is mainly attributed to alterations in age-distribution and population growth [46]. Given the aging trend, the burden of women's cancers might continue to increase in the following decades.

### **Conclusions**

Breast cancer was the most common women's cancer worldwide, which was followed by cervical cancer in the low SDI region. Relatively, ovarian cancer and uterine cancer were more common in high SDI region. Women's cancers cause numerous mortalities worldwide and place heavy economic burdens on women and their families. In the background of global aging, international efforts are needed to reduce women's cancer incidence and mortality and improve women's health.

### Abbreviations

GBD: Global burden disease; DALY: Disability adjusted life year; SDI: Sociodemographic Index; ASIR: Age-standardized incidence rate; ASDR: Agestandardized death rate; EAPC: Estimated annual percentage change

### **Supplementary Information**

The online version contains supplementary material available at https://doi.org/10.1186/s40364-021-00310-y.

Additional file 1: Figure S1: The age standardized rates of breast cancer in 204 countries or territories in 2019. (a) The ASIR of breast cancer in 204 countries or territories; (b) The ASDR of breast cancer in 204 countries or territories; (c) The age-standardized DALY rate of breast cancer in 204 countries or territories. Note: ASIR, Age-standardized incidence rate; ASDR, Age-standardized death rate; DALY, disability adjusted life year.

Additional file 2: Figure S2: The age standardized rates of cervical cancer in 204 countries or territories in 2019. (a) The ASIR of cervical cancer in 204 countries or territories; (b) The ASDR of cervical cancer in 204 countries or territories; (c) The age-standardized DALY rate of cervical cancer in 204 countries or territories. Note: ASIR, Age-standardized incidence rate; ASDR, Age-standardized death rate; DALY, disability adjusted life year.

Additional file 3: Figure S3: The age standardized rates of ovarian cancer in 204 countries or territories in 2019. (a) The ASIR of ovarian cancer in 204 countries or territories; (b) The ASDR of ovarian cancer in 204 countries or territories; (c) The age-standardized DALY rate of ovarian

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cancer in 204 countries or territories. Note: ASIR, Age-standardized incidence rate; ASDR, Age-standardized death rate; DALY, disability adjusted life year.

**Additional file 4: Figure S4:** The age standardized rates of uterine cancer in 204 countries or territories in 2019. (a) The ASIR of uterine cancer in 204 countries or territories; (b) The ASDR of uterine cancer in 204 countries or territories; (c) The age-standardized DALY rate of uterine cancer in 204 countries or territories. Note: ASIR, Age-standardized incidence rate; ASDR, Age-standardized death rate; DALY, disability adjusted life year.

Additional file 5: Figure S5: The incidence, death, DALY, and corresponding age-standardized rates of cancers in different age groups in 2019. (a) The incidence, death, and DALY of cancers in different age groups. (b) The ASIR, ASDR, and age-standardized DALY rate of cancers in different age groups. Note: ASIR, Age-standardized incidence rate; ASDR, Age-standardized death rate; DALY, disability adjusted life year.

**Additional file 6: Figure S6:** The incidence, death, and DALY of cancers in different age groups in high SDI region in 2019. Note: DALY, disability adjusted life year.

**Additional file 7: Figure S7:** The incidence, death, and DALY of cancers in different age groups in high-middle SDI region in 2019. Note: DALY, disability adjusted life year.

**Additional file 8: Figure S8:** The incidence, death, and DALY of cancers in different age groups in middle SDI region in 2019. Note: DALY, disability adjusted life year.

**Additional file 9: Figure S9:** The incidence, death, and DALY of cancers in different age groups in low-middle SDI region in 2019. Note: DALY, disability adjusted life year.

**Additional file 10: Figure S10:** The incidence, death, and DALY of cancers in different age groups in low SDI region in 2019. Note: DALY, disability adjusted life year.

**Additional file 11: Figure S11:** The ASIR, ASDR, and age-standardized DALY rate of cancers in different age groups in high SDI region in 2019. Note: ASIR, Age-standardized incidence rate; ASDR, Age-standardized death rate; DALY, disability adjusted life year.

**Additional file 12: Figure S12:** The ASIR, ASDR, and age-standardized DALY rate of cancers in different age groups in high-middle SDI region in 2019. Note: ASIR, Age-standardized incidence rate; ASDR, Age-standardized death rate; DALY, disability adjusted life year.

**Additional file 13: Figure S13:** The ASIR, ASDR, and age-standardized DALY rate of cancers in different age groups in middle SDI region in 2019. Note: ASIR, Age-standardized incidence rate; ASDR, Age-standardized death rate; DALY, disability adjusted life year.

**Additional file 14: Figure S14:** The ASIR, ASDR, and age-standardized DALY rate of cancers in different age groups in low-middle SDI region in 2019. Note: ASIR, Age-standardized incidence rate; ASDR, Age-standardized death rate; DALY, disability adjusted life year.

**Additional file 15: Figure S15:** The ASIR, ASDR, and age-standardized DALY rate of cancers in different age groups in low SDI region in 2019. Note: ASIR, Age-standardized incidence rate; ASDR, Age-standardized death rate; DALY, disability adjusted life year.

**Additional file 16: Table S1:** The incidence of female breast cancer and temporal trends.

**Additional file 17: Table S2:** The death of female breast cancer and temporal trends.

**Additional file 18: Table S3:** The Disability-Adjusted Life Year (DALY) of female breast cancer and temporal trends.

**Additional file 19: Table S4:** The incidence of female cervical cancer and temporal trends.

**Additional file 20: Table S5:** The death of female cervical cancer and temporal trends.

**Additional file 21: Table S6:** The Disability-Adjusted Life Year (DALY) of female cervical cancer and temporal trends.

Additional file 22:Table S7: The incidence of female ovarian cancer and temporal trends.

**Additional file 23: Table S8:** The death of female ovarian cancer and temporal trends.

**Additional file 24: Table S9:** The Disability-Adjusted Life Year (DALY) of female ovarian cancer and temporal trends.

**Additional file 25: Table S10:** The incidence of uterine cancer and temporal trends.

**Additional file 26: Table S11:** The death of uterine cancer and temporal trends.

**Additional file 27: Table S12:** The Disability-Adjusted Life Year (DALY) of uterine cancer and temporal trends.

**Additional file 28: Table S13:** The age-standardized rate of women's cancers in 204 countries or territories.

### Acknowledgements

Not applicable.

### Authors' contributions

All authors contributed to research performing, drafting and revising the article, gave final approval of the version to be published, and agree to be accountable for all aspects of the work.

### **Funding**

This work was supported by the National Natural Science Foundation of China (No. 82073370 and 81874120).

### Availability of data and materials

All data in this study is available from http://ghdx.healthdata.org/gbd-results-tool.

### Declarations

### Ethics approval and consent to participate

Not applicable.

### Consent for publication

Not applicable.

### Competing interests

The authors declare no conflict of interest.

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## Received: 16 April 2021 Accepted: 20 June 2021 Published online: 07 July 2021

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