



OPEN ACCESS

EDITED BY

Laura Atzori,
University of Cagliari, Italy

REVIEWED BY

Gabriele Biondi,
Azienda Ospedaliero Universitaria Sassari, Italy
Giovanni Deriu,
University of Cagliari, Italy
Claudia Colli,
Dermatological Clinic and STI Center, Italy

*CORRESPONDENCE

Kui Liu
✉ kliu@cdc.zj.cn
Chen Wu
✉ chenwu@cdc.zj.cn

[†]These authors have contributed equally to this work

RECEIVED 12 April 2025

ACCEPTED 13 June 2025

PUBLISHED 07 July 2025

CITATION

Ding Z, Wang S, Li J, Wu H, Lu Q,
Wang X, Fu T, Liu K and Wu C (2025)
Age-period-cohort analysis of syphilis
epidemics in Eastern China, China,
2005–2024.
Front. Public Health 13:1606491.
doi: 10.3389/fpubh.2025.1606491

COPYRIGHT

© 2025 Ding, Wang, Li, Wu, Lu, Wang, Fu, Liu
and Wu. This is an open-access article
distributed under the terms of the [Creative
Commons Attribution License \(CC BY\)](#). The
use, distribution or reproduction in other
forums is permitted, provided the original
author(s) and the copyright owner(s) are
credited and that the original publication in
this journal is cited, in accordance with
accepted academic practice. No use,
distribution or reproduction is permitted
which does not comply with these terms.

Age-period-cohort analysis of syphilis epidemics in Eastern China, China, 2005–2024

Zheyuan Ding^{1†}, Shuangqing Wang^{2†}, Junjie Li¹, Haocheng Wu¹,
Qinbao Lu¹, Xinyi Wang¹, Tianyin Fu¹, Kui Liu^{1*} and Chen Wu^{1*}

¹Department of Public Health Surveillance and Advisory, Zhejiang Provincial Center for Disease Control and Prevention, Hangzhou, Zhejiang Province, China, ²Department of Infectious Disease Control, Quzhou Center for Disease Control and Prevention, Quzhou, Zhejiang Province, China

Background: Syphilis remains one of the serious public health challenges in China and worldwide. This study aims to assess the potential independent risks associated with age, period, and birth cohort for the reported incidence of syphilis in Eastern China.

Methods: Data on all syphilis cases from 2005 to 2024 in Zhejiang Province in Eastern China were collected from the China Information System for Disease Control and Prevention. The Age-Period-Cohort (APC) model was used to analyze the effect coefficients, which were then converted into relative risks (RRs).

Results: From 2005 to 2024, a total of 617,097 syphilis cases were reported in Zhejiang Province. The reported incidence of syphilis decreased by an average of –3.53% per year across all age groups (95% CI: –4.34, –2.70%). The age effect showed that the highest reported incidence was among individuals aged 20–24 years, with a relatively high rate also observed in those aged 60 years and above. The period effect indicated that compared with the reference group of 2010–2014 with the highest reported incidence risk, the risk in 2020–2024 was the lowest (RR = 0.48; 95% CI: 0.43, 0.54). The cohort effect shows that the risk decreased in later birth cohorts, with the highest reported incidence risk in the birth cohort of 1960–1964 (RR = 1.30; 95% CI: 1.08, 1.57) and the lowest risk in the birth cohort of 2020–2024 (RR = 0.00; 95% CI: 0.00, 0.18).

Conclusion: The reported incidence of syphilis in Zhejiang Province has shown an overall downward trend. The implementation of national syphilis control programs has achieved significant results. There is a need to strengthen the management of late-stage syphilis among older adults and enhance syphilis prevention and control efforts among adolescents.

KEYWORDS

syphilis, reported incidence, age-period-cohort model, Eastern China, cohort effect

Introduction

Syphilis is a sexually transmitted infection (STI) caused by *Treponema pallidum* (1). It can also be transmitted from mother to fetus, called congenital syphilis (2). Globally, the number of prevalent syphilis cases was estimated to be 49.71 million in 2019, marking a 60.83% increase from 1990, creating an significant global burden (3). In China, the incidence of syphilis has risen significantly over the past decades, with an average annual growth rate of 11.88% between 2004 and 2019 (4). In June 2010, the Ministry of Health of China launched the National Syphilis Prevention and Control Program (2010–2020) (National Syphilis Control

Program), which aimed to strengthen the prevention and control of syphilis and effectively curb its spread (5). The program set specific control targets for primary and secondary syphilis, as well as congenital syphilis. In 2011, a further national plan was released, specifically targeting the Preventing Mother-to-Child Transmission of HIV, syphilis, and hepatitis B (PMTCT) (6). These programs have achieved significant results. Between 2014 and 2019, the reported incidence of primary and secondary syphilis, as well as congenital syphilis, in China decreased annually by 10.95 and 26.16%, respectively (7). Meanwhile, the incidence risk of syphilis in China varies across different regions. Studies have shown that there is a certain temporal and spatial clustering of syphilis cases in the southeastern coastal areas (8). These regions still face significant challenges in syphilis prevention and control.

As a vital constituent province of the eastern coast of China, Zhejiang Province has a highly developed economy and significant population mobility (9). Over the past decade, Zhejiang Province has consistently ranked among the top in China in terms of the reported incidence of syphilis (4, 7). Moreover, for many years, syphilis has topped the list of notifiable infectious diseases in Zhejiang Province (10), highlighting its significant public health impact. Syphilis prevention and control has always been an important task in the public health domain of Zhejiang Province. Previous studies (10–12) have described the epidemic trends, spatial distribution, and population characteristics of syphilis in Zhejiang Province, but the analysis and exploration of potential causes have been insufficient. Moreover, cross-sectional studies cannot elucidate the independent effects of age, period, and birth cohort on the reported incidence of syphilis. However, the Age-Period-Cohort (APC) model (13) is capable of doing so. But there were few studies on the application of APC in syphilis in China, as well as in Eastern China.

In this study, we chosen Zhejiang Province as the pilot area, to describe the epidemiological characteristics of syphilis in Eastern China, and employed the APC model to analyze the impact of time, age, and birth cohort on the reported incidence trends of syphilis among different genders and different subtypes of syphilis. It would provide the reliable evidence for the available comprehensive action and help to inform more effective public health interventions for further syphilis control.

Methods

Data collection and definitions

Data on syphilis cases in Zhejiang Province from 2005 to 2024 were obtained from the National Notifiable Infectious Disease Reporting Information System, which is a component of the China Information System for Disease Control and Prevention. Syphilis is one of the notifiable infectious diseases in China, and all cases diagnosed by clinicians are required to be reported to this system within 24 h. All

cases included in the study were laboratory-confirmed. The variables included in the analysis were sex, age, occupation, time of onset, and syphilis subtype, which was classified into primary syphilis, secondary syphilis, tertiary syphilis, latent syphilis, and congenital syphilis. All notified syphilis cases were diagnosed and subtyped in accordance with the Chinese National Diagnostic Criteria and management of Syphilis (GB 15974—1995) (14) before 2007, the National Diagnostic Criteria for Syphilis (WS 273—2007) (15) from 2007 to 2017, and the National Diagnostic Criteria for Syphilis (WS 273—2018) after 2018 (16). Primary syphilis, secondary syphilis, and early latent syphilis (infection duration less than 2 years) are classified as early syphilis, while tertiary syphilis and late latent syphilis (infection duration exceeding 2 years) are classified as late syphilis (16, 17). Since latent syphilis is asymptomatic and without signs, it must be detected through screening tests, making it difficult to determine whether it is a new infection or a past infection. Therefore, in this study, the incidence of primary and secondary syphilis is used to reflect the situation of new infections.

Statistical analysis

Epidemiological characteristics of syphilis in Zhejiang Province from 2005 to 2024 were analyzed. APC model was used to evaluate the age, period, and cohort effects of trends in the reported incidence of syphilis among the entire population, as well as separately among males and females. Besides, stratification analyses of APC model were also conducted in primary syphilis, secondary syphilis, tertiary syphilis and latent syphilis. In this study, we divided age into 18 groups, with each group spanning 5 years (0–4, 5–9, ..., 80–84, 85–89). The periods were divided into four intervals (2005–2009, 2010–2014, 2015–2019, and 2020–2024), with the middle period (2010–2014) as the reference. The birth cohorts were determined by subtracting age from the periods, resulting in 21 cohorts (1920–1924, 1925–1929, ..., 2015–2019, 2020–2024), with the middle cohort group (1970–1974) as the reference. The APC model is a type of generalized linear model, with the basic expression as follows:

$$\log R_{ijk} = \log(E_{ij} / N_{ij}) = \mu + \alpha \times \text{Age}_i + \beta \times \text{Period}_j + \gamma \times \text{Cohort}_k + \varepsilon$$

R_{ijk} represents the reported incidence of syphilis in age group i , period j , and birth cohort k ; E_{ij} and N_{ij} represent the expected number of syphilis cases and population at risk in age group i , period j , respectively; μ is the reference level of disease risk; Age_i represents the effect of the age group i ; Period_j represents the effect of the period j ; Cohort_k represents the effect of the birth cohort k associated with the age group i and the period j . Parameters β and γ were converted exponentially to represent the relative risk (RR) of a particular period and birth cohort. The significance of the estimable parameters was evaluated using the two-sided Wald χ^2 test. APC analyses were performed using a web tool from the National Cancer Institute of the United States¹ and R software (version 4.3.0; R Core Team, Vienna, Austria) (13).

Abbreviations: APC, Age-Period-Cohort; CI, Confidence Interval; COVID-19, Coronavirus Disease 2019; EAPC, Estimated Annual Percentage Change; National Syphilis Control Program, National Syphilis Prevention and Control Program (2010–2020); PMTCT, Preventing Mother-to-Child Transmission; RR, Risk Ratio; STD, Sexually Transmitted Disease; STI, Sexually Transmitted Infection.

¹ <https://analysistools.cancer.gov/apc/>

Results

General characteristics of syphilis in Zhejiang Province during study period

Between 2005 and 2024, a total of 617,097 cases of syphilis were reported in Zhejiang Province, including 129,698 cases of primary syphilis, 104,513 cases of secondary syphilis, 3,731 cases of tertiary syphilis, 7,556 cases of congenital syphilis, and 371,598 cases of latent syphilis. The number of female cases was higher than that of male cases. However, for primary syphilis, congenital syphilis, and especially tertiary syphilis, the number of male cases exceeded that of female cases. The majority cases were in the age group of 15–39 years old, accounting for 47.5% of the total cases, followed by the age group of 40–59 years old, which accounted for 31.7%. For tertiary syphilis, the proportion of cases in individuals over 40 years old was relatively higher. Congenital syphilis mainly affected children under 14 years old. Except for congenital syphilis, the other types of syphilis were predominantly found among farmers, who accounted for approximately 40% of the cases (Table 1).

Results of estimated annual percentage change (EAPC) for syphilis

During the study period, the EAPC of the reported incidence of syphilis among all age groups in Zhejiang Province was -3.53% (95% CI: -4.34 , -2.70%). The EAPC varied significantly across different age groups. A downward trend was observed in the 0–9 and 20–49 age

groups, with the notable reductions in the 0–4 and 5–9 age groups, having EAPCs of -29.43% (95% CI: -45.19 , -9.14%) and -16.94% (95% CI: -25.95 , -6.83%), respectively. Additionally, the 25–29, 30–34, and 35–39 age groups also experienced significant declines, with EAPCs of -8.81% (95% CI: -10.00 , -7.61%), -8.61% (95% CI: -9.74 , -7.46%), and -7.23% (95% CI, -8.42 , -6.03%), respectively. However, the 15–19 age group showed an upward trend, with an EAPC of 3.68% (95% CI, 1.14 , 6.28%) (Figure 1).

The reported incidence of syphilis declined by an average of 3.00% (95% CI: -3.78 , -2.21%) per year among males and 3.08% (95% CI: -4.07 , -2.07%) per year among females. Among the 0–9 and 35–49 age groups, the decline was greater in males than in females, while among the 20–34 age group, the decline was greater in females than in males (Supplementary Figure S1).

The further stratified analysis showed that the reported incidence of primary and secondary syphilis declined across all age groups. The EAPCs were -14.67% (95% CI: -17.97 , -11.23%) for primary syphilis and -8.43% (95% CI: -9.82 , -7.03%) for secondary syphilis, with a greater decline observed in females than in males. In contrast, latent syphilis incidence increased slightly, with an EAPC of 1.97% (95% CI: 0.80 , 3.16%), which were higher in males than in females. The EAPC for tertiary syphilis was not statistically significant. Among individuals aged ≥ 15 years, the reported incidence of primary syphilis decreased across all age groups, with a remarkable decline in females. For secondary syphilis, the reported incidence decreased across all age groups of individuals aged ≥ 20 years, with a greater decline in females aged 20–49 years. The reported incidence of tertiary syphilis increased in individuals aged ≥ 70 years. For latent syphilis, the reported

TABLE 1 Epidemiological characteristics of syphilis in Zhejiang Province from 2005 to 2024.

Variables	Syphilis	Primary syphilis	Secondary syphilis	Tertiary syphilis	Congenital syphilis	Latent syphilis
Total, <i>n</i> (%)	617,097(100)	129,698(21.0)	104,513(16.9)	3,731(0.6)	7,556(1.2)	371,598(60.2)
Sex, <i>n</i> (%)						
Male	288,740(46.8)	66,629(51.4)	51,156(48.9)	2,593(69.5)	4,162(55.1)	164,199(44.2)
Female	328,357(53.2)	63,069(48.6)	53,357(51.1)	1,138(30.5)	3,394(44.9)	207,399(55.8)
Age group (years), <i>n</i> (%)						
0–14	8,662(1.4)	347(0.3)	211(0.2)	4(0.1)	7,515(99.5)	585(0.2)
15–39	293,216(47.5)	73,659(56.8)	60,826(58.2)	792(21.2)	30(0.4)	157,909(42.5)
40–59	195,528(31.7)	40,352(31.1)	33,189(31.8)	1,628(43.6)	4(0.1)	120,355(32.4)
≥ 60	119,690(19.4)	15,340(11.8)	10,287(9.8)	1,307(35.0)	7(0.1)	92,749(25.0)
Population classification, <i>n</i> (%)						
Farmer	247,378(40.1)	51,945(40.1)	38,954(37.3)	1,531(41.0)	23(0.3)	154,925(41.7)
Houseworkers or unemployed	100,955(16.4)	17,128(13.2)	15,420(14.8)	598(16.0)	9(0.1)	67,800(18.2)
Industrial workers	62,960(10.2)	16,881(13.0)	14,259(13.6)	336(9.0)	3(0.0)	31,481(8.5)
Commercial service worker	51,669(8.4)	9,805(7.6)	9,666(9.2)	292(7.8)	6(0.1)	31,900(8.6)
Retiree	24,754(4.0)	2,687(2.1)	1910(1.8)	308(8.3)	4(0.1)	19,845(5.3)
Migrant worker	20,855(3.4)	8,114(6.3)	4,512(4.3)	83(2.2)	2(0.0)	8,144(2.2)
Preschool children	7,613(1.2)	261(0.2)	104(0.1)	2(0.1)	6,900(91.3)	346(0.1)
Others	100,912(16.4)	22,877(17.6)	19,688(18.8)	581(15.6)	609(8.1)	57,157(15.4)

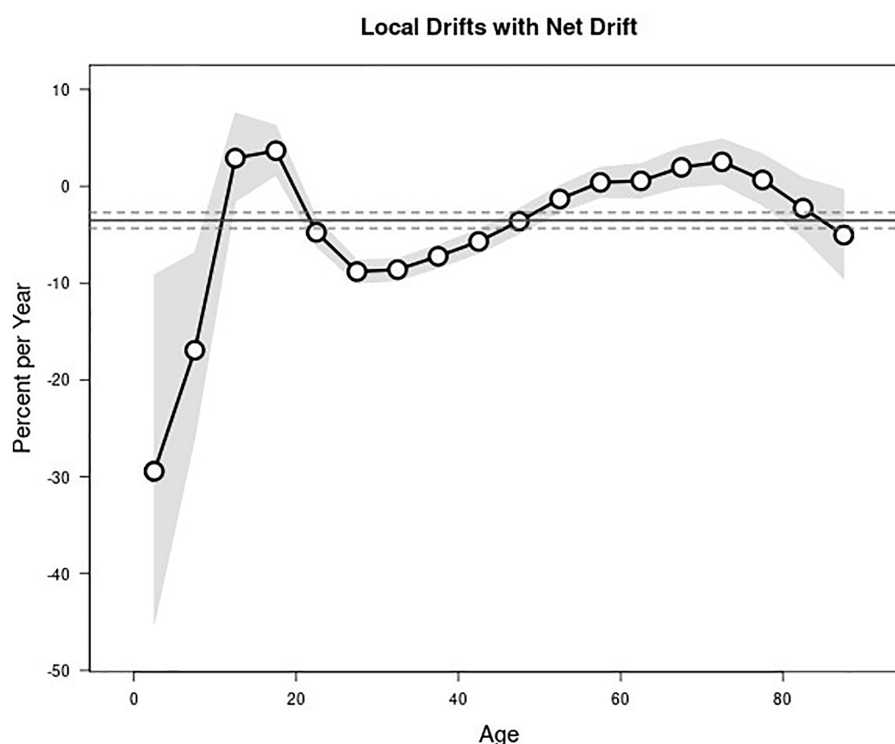


FIGURE 1
Estimated annual percentage change of reported incidence of syphilis in each age groups in Zhejiang province.

incidence increased in individuals aged 10–14 years (EAPC: 12.36, 95% CI: 1.98, 23.79%) and 15–19 years (EAPC: 9.91, 95% CI: 7.25, 12.63%), and showed a slight increase in those aged 45–79 years, while it decreased in individuals aged 25–39 years (Supplementary Tables S1–S4).

Age effect

Given that the absolute value of the net drift exceeded 1%, longitudinal age curves were utilized in this study to delineate the age-specific effects. After adjusting for period and birth cohort influences, the longitudinal age curves revealed distinct patterns in the reported incidence across different age groups. Specifically, the 0–4 age group exhibited a relatively high incidence, which then declined sharply in both the 5–9 and 10–14 age groups, resulting in the lowest reported incidence among these cohorts. After the age of 15, the reported incidence began to rise significantly, peaking in the 20–24 age group. Subsequently, the reported incidence decreased with increasing age, although a slight rebound was observed after the age of 60 (Figure 2).

The age effects in males and females were basically consistent with the overall population effect. It is worth noting that the age group with the highest reported incidence in males was the 25–29 years group, while in females it was the 20–24 years group. Among individuals aged 55–69 years, the reported incidence in males was higher than that in females of the same age group, while in other age groups, the incidence in males was lower than that in females (Supplementary Figures S2, S3).

For primary and secondary syphilis, the reported incidences were relatively higher among individuals aged 15–34 years, with the 20–24 years group having the highest reported incidence. For tertiary syphilis, the reported incidence was relatively higher among individuals aged 80 years and above. For latent syphilis, there was a small peak in the reported incidence among individuals aged 20–34 years, followed by a slight decline, and then a significant increase from the age of 60 years, with the highest rate observed in the 85–89 years group. Overall, in all stages of syphilis, the reported incidence was higher in females than in males among younger age groups, while among older age groups, the reported incidence was higher in males than in females (Supplementary Tables S5–S8).

Period effect

After controlling for age and cohort variables, the risk on reported incidence of syphilis increased from 2005–2009 to 2010–2014 and then decreased afterward. The lowest risk was observed during 2020–2024 (RR = 0.48; 95% CI: 0.43, 0.54; $p < 0.001$). The analysis results for males and females were generally consistent with the overall population effect (Figure 2 and Supplementary Figures S2, S3).

From 2005–2009 to 2010–2014, the risk on reported incidence of secondary syphilis increased. The risk on reported incidence of primary and secondary syphilis decreased after 2010–2014. For latent syphilis, the risk increased from 2005–2009 to 2015–2019 and then decreased afterward. The period effect for tertiary syphilis was not statistically significant. The results for males and females were generally consistent (Supplementary Tables S5–S8).

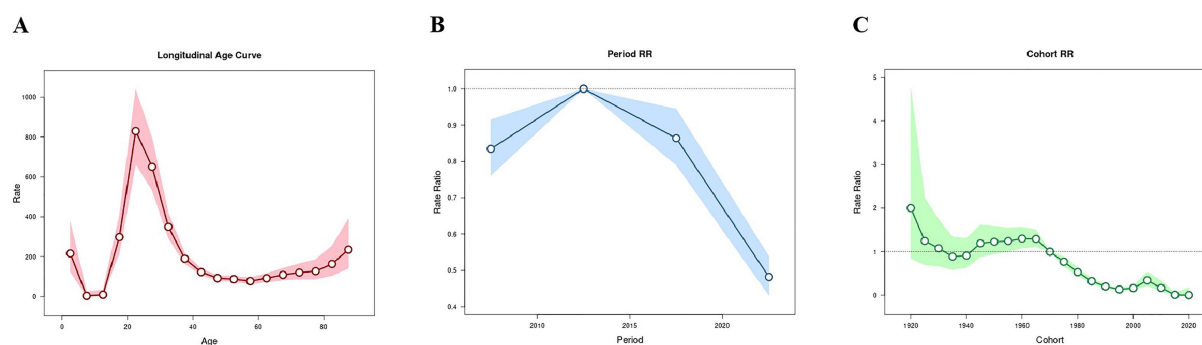


FIGURE 2

Age-period-cohort effects on the reported incidence of syphilis in Zhejiang Province. (A) Age effect. (B) Period effect. (C) Cohort effect. The colored area represented the 95% confidence interval (CI).

Cohort effect

Overall, the risk on reported incidence of syphilis decreased progressively with more recent birth cohorts. Using the 1970–1974 birth cohort as the reference, cohorts born between 1955 and 1969 exhibited higher risks of syphilis, with the highest risk observed in 1960–1964 birth cohort (RR = 1.30; 95% CI: 1.08, 1.57). In contrast, cohorts born after 1975 had a risk less than 1 with a successive declined trend. However, a slight increase was noted in the cohorts born in 2000–2004 and 2005–2009. The lowest risk was in the 2020–2024 cohort (RR = 0.00; 95% CI: 0.00, 0.18). Cohorts born between 1920 and 1954 showed no statistically significant differences. The findings for males and females were largely consistent with the overall population trends (Figure 2 and Supplementary Figures S2, S3).

For primary syphilis, the risk on reported incidence decreased with more recent birth cohorts. The highest risk was observed in the 1920–1924 cohort, while the lowest risk was in the 2015–2019 cohort. For secondary syphilis, a downward trend was also observed. The highest risk was in the 1920–1924 cohort, and the lowest risk was in the 1995–1999 cohort. For tertiary syphilis, cohorts born between 1920 and 1939 had a risk lower than that of the reference cohort (1970–1974), while other cohorts showed no statistically significant differences. For latent syphilis, cohorts born between 1920 and 1964 and between 1985 and 1999 had a risk lower than the reference cohort, while the cohort born in 2005–2009 had a higher risk than the reference cohort. The findings for males and females were generally consistent with the overall population trends (Supplementary Tables S5–S8).

Discussion

This study conducted a systematic analysis of the reported incidence of syphilis in Zhejiang Province, an eastern coastal region of China, using the APC model. It explored the independent risks associated with age, period, and birth cohort, as well as the potential influencing of factors such as age, gender, socioeconomic changes, syphilis prevention policies, sexual attitudes, and the Coronavirus disease 2019 (COVID-19) pandemic. These findings are expected to play a critical role in promoting the effective implementation of syphilis interventions, thereby reducing the disease burden of syphilis in Eastern China.

Over the past two decades, the reported incidence of syphilis in Zhejiang Province has shown a general downward trend. This differs from the epidemiological trends of syphilis in other countries. Japan has experienced a significant increase in syphilis cases after 2011 (18, 19), while South Korea has seen a more moderate rise (19). The United States also observed an increase in syphilis incidence between 2017 and 2024 (20). In Canada, the incidence of infectious syphilis began to rise steadily in the early 2000s and surged sharply after 2017 (21). These differences may be related to variations in sexual attitudes and behaviors, healthcare accessibility, and public health policies for syphilis among different countries.

Since 2010–2014, there has been a significant reduction in the reported incidence of primary and secondary syphilis in Zhejiang Province. The reported incidence in the 0–9 age group has also declined rapidly over these 20 years, which largely due to the effective control of congenital syphilis. These results implied a downward trend in new syphilis infections and cases of congenital syphilis, which was largely attributed to the successful implementation of the National Syphilis Control Program (5) and PMTCT (6). The effectiveness of these programs has also been proved in other southeastern area in China (22). The specific measures of these two programs include providing universally accessible and free syphilis counseling services, enhancing screening program to promote early diagnosis and treatment, offering standardized, high-quality and accessible diagnostic and treatment services, carrying out extensive health education campaigns, promoting condom use, as well as early detection and intervention for syphilis among pregnant women. Meanwhile, the latent syphilis had shown a slight overall upward trend. Specifically, there was a continuous increase before 2015–2019, followed by a decline from 2020 to 2024. The earlier rise might be mainly due to the standardization of testing and reporting under the National Syphilis Control Program. It encompassed the enhancement of the national syphilis surveillance network, the strengthening of quality control for syphilis testing, and the implementation of active syphilis screening. The subsequent decline was likely attributable to the impact of the COVID-19 pandemic. Although some countries saw a rise in syphilis cases during or after this period (23, 24), in China, STIs such as HIV, gonorrhea, syphilis, hepatitis B, and hepatitis C all presented a decline (25–27). Available evidence demonstrated that the number of reported syphilis cases in China decreased by 17.95% in 2020 and 20.41% in 2021, respectively (28). This phenomenon was not due to decreased

diagnostic accuracy or underreporting of syphilis (28). Instead, the pandemic led to a reallocation of medical resources toward COVID-19 control, resulting in weakened or temporarily interrupted routine HIV/STI screening, diagnostic, and treatment services (29). Additionally, restrictions on population mobility were likely to reduce the healthcare-seeking behavior of asymptomatic individuals or cases with mild symptoms that experienced high risk of exposure (30).

The number of reported syphilis cases among sexually active individuals aged 20–34 years was relatively high, especially for primary and secondary syphilis, which have higher incidence in this population. Compared to males, females had a higher risk, and the peak age group was younger. This might be due to differences between males and females in biological risk of STI infection, clinical manifestations, and sexual behavior characteristics (31), causing STIs more easily transmitted from males to females (32). Moreover, this specific age group was the childbearing period for females, and prenatal screening increased the detection of syphilis. The reported incidence of syphilis in the 15–19 age group showed an upward trend over the past 20 years, with a rapid increase in the reported incidence of latent syphilis in the 10–19 age group. This indicated that syphilis infection in Zhejiang Province was beginning to affect younger populations. This is consistent with the increasing trend of HIV infections among young students in China (33). This might be because contemporary youth were more physiologically and psychologically precocious, with related liberal sexual attitudes (34). Additionally, China tended to adopt abstinence-based education strategies, which could result in adolescents with insufficient knowledge about STIs (35), lack of self-protection awareness, and a higher likelihood of engaging in risky sexual behaviors (36). Thus, comprehensive sex education programs should be introduced for young people, along with support from families and society, as well as sexual and reproductive health services, to reduce the risk of syphilis transmission (34).

Tertiary syphilis was associated with severe complications affecting the nervous system, cardiovascular system, eyes, and other organs (37), which could significantly impair quality of life. Meanwhile, latent syphilis, with its large case base and comprising over half of all syphilis cases, also contributed substantially to the overall disease burden. This study observed an increase in tertiary syphilis among individuals aged 70 and above. Moreover, the reported incidence of syphilis, particularly tertiary and latent syphilis, were found to be higher in the older adult population. These findings were consistent with other studies in Guangdong Province (38), Shandong Province (39), and Japan (40). Among older adults, especially in rural areas, commercial sex and high-risk sexual behaviors were relatively common while condom use was infrequent (41, 42). Compared to younger individuals, older adults generally had lower awareness of the risks associated with STIs and experienced higher levels of stigma (38). Besides, physiological changes during aging, such as increased fragility of the vaginal mucosa and micro-abrasions after intercourse, could elevate the risk of STIs (43). Additionally, older adults often pay less attention to infections, leading to delayed medical visits (44) and interrupt treatment. This frequently results in early-stage syphilis progressing to tertiary syphilis. Besides, given that older adults were more likely than younger individuals to be hospitalized for underlying diseases, the higher reported incidence of latent syphilis among older adults may be attributed to the increased likelihood of syphilis screening and detection during hospitalization. Therefore, it was a need to

strengthen sexual health education among older adults, disseminate knowledge about STIs, advocate the use of condoms, and eliminate feelings of shame and inferiority. Simultaneously, sufficient medical services and effective health support should be provided for older adults, including the establishment of geriatric sexual health clinics, and STI screening for high-risk older adult groups to reduce the risk of syphilis and other STIs among the older adult population.

As birth cohorts progressed, the reported risk of syphilis, particularly for primary and secondary syphilis, declined. This trend was largely attributed to a series of syphilis control campaigns in China. In the nascent years of New China, following its founding in 1949, syphilis emerged as one of the most pressing public health challenges (45). Between the early 1950s and 1964, China initiated its first comprehensive nationwide campaign against syphilis (46). This multifaceted effort encompassed the closure of brothels, the implementation of extensive health education programs, widespread screening initiatives, and the provision of free treatment for those infected. By 1964, these concerted actions had basically eradicated syphilis across the nation (46, 47). Subsequently, syphilis virtually disappeared from public view. However, in the early 1980s, with the comprehensive implementation of economic reform and opening-up policies, significant changes occurred in the socio-economic and life environment, as well as in people's attitudes toward sex. These changes included increased economic levels, greater population mobility, commercial sex, and extramarital sex, all of which contributed to the resurgence of syphilis (48, 49). In response, China launched its second nationwide syphilis control campaign (49), including establishing the Chinese National Center for STD (Sexually Transmitted Disease) Control in 1986 and the national STD surveillance system in 1987. Health education and behavior change initiatives, with a focus on promoting condom use, have been prioritized as key public health strategies to address the syphilis epidemic (46). Since the late 1990s, the annual growth rate of syphilis has slowed down but remains high (50), particularly with increasing prevalence among high-risk populations such as female sex workers and men who have sex with men (51, 52). According to our findings, cohorts born before 1970–1974 exhibited a continuous increase in the reported risk of latent syphilis. Additionally, the reported incidence of tertiary syphilis in the 1970–1974 birth cohort was higher than that in cohorts born before 1935–1939. These patterns might be attributed to the impact of these historical junctures, as well as the development of the internet, which has facilitated the use of social software to seek sexual partners and thereby increased the difficulty of syphilis prevention and control. The study also observed a slight increase in cohorts born in 2000–2004 and 2005–2009, likely due to the intensified syphilis testing efforts under the National Syphilis Control Program implemented after 2010.

Limitations

Some limitations also should be mentioned. First, our data was derived from a passive surveillance network, which might be subject to reporting bias. Secondly, this study lacked a specific analysis of congenital syphilis. Since the potential population affected by congenital syphilis were neonates rather than the general population, it was not feasible to conduct relevant analyses using the APC model. Third, as an emerging intervention for syphilis prevention, pre-exposure prophylaxis for HIV (PrEP) was not thoroughly discussed in this study due to the lack of patient-related data.

Conclusion

Over the past 20 years, the reported incidence of syphilis in Zhejiang Province showed a general downward trend. Individuals aged 20–34 years were the most susceptible population. However, the high incidence of late-stage syphilis among older adults and the rising risk of syphilis among adolescents were serious issues that warranted attention. These findings could help relevant departments through formulating appropriate policies and implementing targeted prevention and control measures in susceptible individuals.

Data availability statement

The original contributions presented in the study are included in the article/[Supplementary material](#), further inquiries can be directed to the corresponding author.

Ethics statement

The studies involving humans were approved by Zhejiang Provincial Center for Disease Control and Prevention. The studies were conducted in accordance with the local legislation and institutional requirements. Written informed consent for participation was not required from the participants or the participants' legal guardians/next of kin in accordance with the national legislation and institutional requirements.

Author contributions

ZD: Writing – original draft, Conceptualization. SW: Writing – original draft, Validation, Supervision. JL: Validation, Formal analysis, Writing – review & editing, Methodology. HW: Writing – review & editing, Funding acquisition. QL: Data curation, Writing – review & editing. XW: Data curation, Validation, Writing – review & editing, Supervision. TF: Validation, Supervision, Writing – review & editing, Resources, Formal analysis. KL: Writing – original draft, Conceptualization, Funding acquisition, Supervision. CW: Conceptualization, Validation, Supervision, Writing – review & editing.

References

1. Zhou S, Chanderraj R. What is syphilis? *JAMA*. (2023) 329:1710–0. doi: 10.1001/jama.2023.2897
2. Sankaran D, Partridge E, Lakshminrusimha S. Congenital syphilis—an illustrative review. *Children*. (2023) 10:1310. doi: 10.3390/children10081310
3. Chen T, Wan B, Wang M, Lin S, Wu Y, Huang J. Evaluating the global, regional, and national impact of syphilis: results from the global burden of disease study 2019. *Sci Rep*. (2023) 13:11386. doi: 10.1038/s41598-023-38294-4
4. Li J, Yang Y, Huang B, Zeng J. Epidemiological characteristics of syphilis in mainland China, 2004 to 2019. *J Int Med Res*. (2024) 52:3000605241258465. doi: 10.1177/03000605241258465
5. China Ministry of Health. Notice of the Ministry of Health on issuing National Program for prevention and control of syphilis in China (2010–2020). Available online at: http://www.gov.cn/gzdt/2010-06/21/content_1632301.htm (accessed March 1, 2025)
6. China Ministry of Health. Notice of the ministry of health on issuing the implementation plan for preventing mother-to-child transmission of HIV, syphilis, and hepatitis B. Available online at: <https://www.nhc.gov.cn/wjw/c100175/201102/8d21517b4b2e487bb4192136f0fea00f.shtml> (Accessed June 23, 2025)
7. Yue X, Gong X, Li J, Zhang J. Epidemiological trends and features of syphilis in China, 2014–2019. *Chin J Dermatol*. (2021) 54:668–72. doi: 10.35541/cjd.20210098
8. Chen Z, Liang S, Yue X, Li J, Zhang J, Gong X. Epidemiological trends and spatio-temporal distribution characteristics of syphilis in China from 2010 to 2023. *Chin J Dermatol*. (2024) 57:1045–50. doi: 10.35541/cjd.20240073
9. Liu K, Chen S, Zhang Y, Li T, Xie B, Wang W, et al. Tuberculosis burden caused by migrant population in eastern China: evidence from notification records in Zhejiang Province during 2013–2017. *BMC Infect Dis*. (2022) 22:109. doi: 10.1186/s12879-022-07071-5

Funding

The author(s) declare that financial support was received for the research and/or publication of this article. This study was supported by the Zhejiang Provincial Medical and Health Project (WKJ-ZJ-2522, 2024KY895, and 2025KY774).

Conflict of interest

The authors declare that the research was conducted in the absence of any commercial or financial relationships that could be construed as a potential conflict of interest.

Generative AI statement

The authors declare that no Gen AI was used in the creation of this manuscript.

Publisher's note

All claims expressed in this article are solely those of the authors and do not necessarily represent those of their affiliated organizations, or those of the publisher, the editors and the reviewers. Any product that may be evaluated in this article, or claim that may be made by its manufacturer, is not guaranteed or endorsed by the publisher.

Supplementary material

The Supplementary material for this article can be found online at: <https://www.frontiersin.org/articles/10.3389/fpubh.2025.1606491/full#supplementary-material>

SUPPLEMENTARY FIGURE S1

Estimated annual percentage change of reported incidence of syphilis in each age groups and gender in Zhejiang Province.

SUPPLEMENTARY FIGURE S2

Age-period-cohort effects on the reported incidence of syphilis in males in Zhejiang Province.

SUPPLEMENTARY FIGURE S3

Age-period-cohort effects on the reported incidence of syphilis in females in Zhejiang Province.

10. Wu H, Xue M, Wu C, Lu Q, Ding Z, Wang X, et al. Scaling law characteristics and spatiotemporal multicomponent analysis of syphilis from 2016 to 2022 in Zhejiang Province, China. *Front Public Health*. (2023) 11:1275551. doi: 10.3389/fpubh.2023.1275551
11. Zhu X, Zhu Z, Gu L, Zhan Y, Gu H, Yao Q, et al. Spatio-temporal variation on syphilis from 2005 to 2018 in Zhejiang Province, China. *Front Public Health*. (2022) 10:873754. doi: 10.3389/fpubh.2022.873754
12. Yao Q, Zeng F, Fei L, Kong W, Du N, Wu L, et al. Epidemiology of syphilis in Zhejiang province, 2010–2019. *Chin J Epidemiol*. (2020) 41:1313–8. doi: 10.3760/cma.j.cn112338-20200324-00425
13. Rosenberg PS, Check DP, Anderson WF. A web tool for age-period-cohort analysis of cancer incidence and mortality rates. *Cancer Epidemiol Biomarkers Prev*. (2014) 23:2296–302. doi: 10.1158/1055-9965.epi-14-0300
14. China Ministry of Health (1995) Chinese National Diagnostic Criteria and management of syphilis (GB 15974–1995)
15. China Ministry of Health National diagnostic criteria for syphilis (WS 273–2007) (2007)
16. China National Health and Family Planning Commission (2018) National diagnostic criteria for syphilis (WS 273–2018)
17. World Health Organization. WHO guidelines approved by the guidelines review committee. WHO guidelines for the treatment of *Treponema pallidum* (syphilis). Geneva: World Health Organization (2016).
18. Mori Y, Miyatake N, Mori Y, Tanimoto K, Morioka H. Trends in syphilis incidence and its association with the number of sex industry-related businesses in Japan: an ecological study using Joinpoint analysis. *Cureus*. (2025) 17:e83294. doi: 10.7759/cureus.83294
19. Lee M-J, Komatsu S. Epidemiological trends of syphilis infections in Japan and South Korea from 2011 to 2019. *Clin Infect Immun*. (2025) 10:9–17. doi: 10.14740/cii181
20. Do D, Rodriguez PJ, Gratzl S, Cartwright BMG, Baker C, Stucky NL. Trends in incidence of syphilis among US adults from January 2017 to October 2024. *Am J Prev Med*. (2025) 2:8. doi: 10.1016/j.amepre.2025.03.008
21. Aho J, Lybeck C, Tetteh A, Issa C, Kouyoumdjian F, Wong J, et al. Rising syphilis rates in Canada, 2011–2020. *Can Commun Dis Rep*. (2022) 48:52–60. doi: 10.14745/ccdr.v48i23a01
22. Shi L, Chen L, Liu X, Hu H, Chen Y, Chen Y, et al. Evaluating the effect of the plan of national syphilis control in controlling the syphilis epidemic in Jiangsu, China 2010–2020. *Front Public Health*. (2023) 11:1281229. doi: 10.3389/fpubh.2023.1281229
23. Tsantes AG, Tournas P, Domouchtsidou A, Nicolaidou E, Bonovas S, Stratigos A, et al. The resurgence of *Treponema pallidum* infections and reinfections during the COVID-19 pandemic in Greece. *Int J Environ Res Public Health*. (2024) 21:1283. doi: 10.3390/ijerph21101283
24. Komori A, Mori H, Xie W, Valenti S, Naito T. Rapid resurgence of syphilis in Japan after the COVID-19 pandemic: a descriptive study. *PLoS One*. (2024) 19:e0298288. doi: 10.1371/journal.pone.0298288
25. Wu X, Zhou X, Chen Y, Zhai K, Sun R, Luo G, et al. The impact of COVID-19 lockdown on cases of and deaths from AIDS, gonorrhea, syphilis, hepatitis B, and hepatitis C: interrupted time series analysis. *JMIR Public Health Surveill*. (2023) 9:e40591. doi: 10.2196/40591
26. Yan J, Li Y, Zhou P. Impact of COVID-19 pandemic on the epidemiology of STDs in China: based on the GM (1,1) model. *BMC Infect Dis*. (2022) 22:519. doi: 10.1186/s12879-022-07496-y
27. Xu A, Zuo Z, Yang C, Ye F, Wang M, Wu J, et al. A long trend of sexually transmitted diseases before and after the COVID-19 pandemic in China (2010–21). *Sex Health*. (2023) 20:497–505. doi: 10.1071/sh22172
28. Wu Y, Zhu W, Yue X, Li J, Zhang J, Gong X, et al. Impact of COVID-19 epidemic on syphilis case reporting in China. *Chin J Epidemiol*. (2022) 43:2015–20. doi: 10.3760/cma.j.cn112338-20220920-00794
29. Liao W. The impact on prevention and control strategies for HIV/STIs from COVID-19 pandemic to post-pandemic period in southern China (2020–2023): a literature review. *Theoret Nat Sci*. (2024) 59:246–58. doi: 10.54254/2753-8818/59/20241424
30. Sentís A, Prats-Urbe A, López-Corbeto E, Montoro-Fernandez M, Nomah DK, de Olalla PG, et al. The impact of the COVID-19 pandemic on sexually transmitted infections surveillance data: incidence drop or artefact? *BMC Public Health*. (2021) 21:1637. doi: 10.1186/s12889-021-11630-x
31. Hook EW. Gender differences in risk for sexually transmitted diseases. *Am J Med Sci*. (2012) 343:10–1. doi: 10.1097/MAJ.0b013e31823ea276
32. Wong T, Singh A, Mann J, Hansen L, McMahon S. Gender differences in bacterial STIs in Canada. *BMC Womens Health*. (2004) 4:10–11. doi: 10.1186/1472-6874-4-s1-s26
33. Cai C, Tang H, Chen F, Li D, Lyu F. Characteristics and trends of newly reported HIV infection in young students in China, 2010–2019. *Chin J Epidemiol*. (2020) 41:1455–9. doi: 10.3760/cma.j.cn112338-20200417-00592
34. Yu J. An overview of the sexual behaviour of adolescents and young people in contemporary China. *AMJ*. (2010) 3:397–403. doi: 10.4066/AMJ.2010.317
35. Li C, Cheng Z, Wu T, Liang X, Gaoshan J, Li L, et al. The relationships of school-based sexuality education, sexual knowledge and sexual behaviors—a study of 18,000 Chinese college students. *Reprod Health*. (2017) 14:103. doi: 10.1186/s12978-017-0368-4
36. Lyu J, Shen X, Hesketh T. Sexual knowledge, attitudes and Behaviours among undergraduate students in China-implications for sex education. *Int J Environ Res Public Health*. (2020) 17:6716. doi: 10.3390/ijerph17186716
37. Ghanem KG, Ram S, Rice PA. The modern epidemic of syphilis. *N Engl J Med*. (2020) 382:845–54. doi: 10.1056/NEJMra1901593
38. Wang C, Zhao P, Xiong M, Tucker JD, Ong JJ, Hall BJ, et al. New syphilis cases in older adults, 2004–2019: an analysis of surveillance data from South China. *Front Med*. (2021) 8:781759. doi: 10.3389/fmed.2021.781759
39. Liu D, Yang Y, Xiu C, Li Z, Chu T, Tian H. Trends in the prevalence of syphilis among older adults in Shandong, China. *Southeast Asian J Trop Med Public Health*. (2016) 47:1192–7.
40. Takahashi M, Hagiya H, Koyama T, Otsuka F. Trends in the incidence of syphilis in the middle-aged and older adults in Japan: a nationwide observational study, 2009–2019. *Geriatr Gerontol Int*. (2022) 22:1019–24. doi: 10.1111/ggi.14500
41. Li Y, Liu Q, Yang Y, Fan S, Liu Y, Li N. A study on the association between family support and high-risk sexual behavior of elderly men in rural China. *Am J Mens Health*. (2022) 16:15579883221107729. doi: 10.1177/15579883221107729
42. Li N, Li H, Ma Y, Fan P, Yang W, Zhu Q, et al. A qualitative study on high risk behaviors and related factors of reported HIV/AIDS cases aged 60 years and above in some areas of Henan province. *Chin J Epidemiol*. (2017) 38:1161–4. doi: 10.3760/cma.j.issn.0254-6450.2017.09.003
43. Johnson BK. Sexually transmitted infections and older adults. *J Gerontol Nurs*. (2013) 39:53–60. doi: 10.3928/00989134-20130918-01
44. Wong NS, Huang S, Zheng H, Chen L, Zhao P, Tucker JD, et al. Stages of syphilis in South China – a multilevel analysis of early diagnosis. *BMC Public Health*. (2017) 17:135. doi: 10.1186/s12889-016-4004-y
45. Cohen MS, Ping G, Fox K, Henderson GE. Sexually transmitted diseases in the People's Republic of China in Y2K: back to the future. *Sex Transm Dis*. (2000) 27:143–5. doi: 10.1097/00007435-200003000-00004
46. Chen XS, Yin YP, Wang QQ, Wang BX. Historical perspective of syphilis in the past 60 years in China: eliminated, forgotten, on the return. *Chin Med J*. (2013) 126:2774–9. doi: 10.3760/cma.j.issn.0366-6999.20123559
47. Cohen MS, Henderson GE, Aiello P, Zheng H. Successful eradication of sexually transmitted diseases in the People's Republic of China: implications for the 21st century. *J Infect Dis*. (1996) 174:S223–9. doi: 10.1093/infdis/174.supplement_2.s223
48. Abrams HK. The resurgence of sexually transmitted disease in China. *J Public Health Policy*. (2001) 22:429–40. doi: 10.2307/3343160
49. Chen X-S, Jiang T-T, Yin Y-P, Wang Q-Q. The past 70 years in control of syphilis in China: elimination and responses to resurgence. *Int J Dermatol Venereol*. (2020) 3:193–7. doi: 10.1097/JD9.0000000000000134
50. Chen ZQ, Zhang GC, Gong XD, Lin C, Gao X, Liang GJ, et al. Syphilis in China: results of a national surveillance programme. *Lancet*. (2007) 369:132–8. doi: 10.1016/s0140-6736(07)60074-9
51. Lin CC, Gao X, Chen XS, Chen Q, Cohen MS. China's syphilis epidemic: a systematic review of seroprevalence studies. *Sex Transm Dis*. (2006) 33:726–36. doi: 10.1097/01.olq.0000222703.12018.58
52. Tucker JD, Yin YP, Wang B, Chen XS, Cohen MS. An expanding syphilis epidemic in China: epidemiology, behavioural risk and control strategies with a focus on low-tier female sex workers and men who have sex with men. *Sex Transm Infect*. (2011) 87:ii16–8. doi: 10.1136/sti.2010.048314