Addiction

HIV epidemic among drug users in China: 1995-2011

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ABSTRACT

Aim To describe trends in the HIV epidemic among drug users (DUs) in China from 1995 to 2011. **Design, Setting and Participants** Data sets from China's national HIV/AIDS case reporting and sentinel surveillance systems as of December 2011 were used separately for descriptive analysis. **Measures** Changes in the geographic distribution of the number of HIV cases and HIV prevalence among injecting drug users (IDUs) and non-IDUs were examined. We also analysed changes in HIV prevalence among the broader DU population, and drug use-related behaviours including types of drugs used, recent injecting and recent needle sharing in the context of the rapid scale-up of DU sentinel sites and national harm reduction programmes. **Findings** The HIV epidemic among China's DUs is still highly concentrated in five provinces. Here, HIV prevalence peaked at 30.3% [95% confidence interval (CI) = 28.6, 32.1] among IDUs in 1999, and then gradually decreased to 10.9% (95% CI = 10.6, 11.2) by 2011. We observed a rapid increase in the use of 'nightclub drugs' among DUs from 1.3% in 2004 to 24.4% in 2011. A decline in recent needle sharing among current IDU from 19.5% (95% CI = 19.4, 19.6) in 2006 to 11.3% (95% CI = 11.2, 11.4) in 2011 was found to be correlated with the rapid scale-up of methadone maintenance treatment (MMT; $r_{(4)} = -0.94$, P = 0.003) harm reduction efforts. **Conclusions** While HIV prevalence and needle sharing among current injecting drug users in China have declined dramatically and are correlated with the scale-up of national harm reduction efforts, the recent, rapid increased use of 'nightclub drugs' presents a new challenge.

Keywords Behaviour, China, drug user, harm reduction, HIV infection, epidemic trend.

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INTRODUCTION

The earliest evidence of an HIV epidemic in China was identified in 1989, among injecting drug users (IDUs) in southern Yunnan Province [1]. Findings from geospatial analysis of HIV subtypes, behavioural surveys among IDUs and their sex partners and routine surveillance activities indicate that needle sharing among IDUs, and sexual contact between IDUs and sex workers, were the major driver of later subepidemics across mainland China [2]. At the end of 2011, the number of people infected with HIV via injecting drug use was estimated at some 221 000 [3].

In 1985 the Chinese government had established a national HIV case reporting system (CRS) [4,5], when the first ever case of HIV on the mainland was identified

in a foreign visitor [4]. From 1985 until 1995, CRS remained China's primary surveillance tool. In 1995, the government launched the sentinel surveillance system to conduct cross-sectional surveys in 42 sites among four key affected populations (KAPs)-drug users (DUs, including those who inject and those who do not), female sex workers, sexually transmitted disease clinic patients and long-distance truck drivers. This system expanded and evolved in 2004 to monitor both HIV prevalence and HIV related behaviours [6,7] among HIV KAPs. By 2010, the total number of HIV sentinel surveillance sites had been expanded to 1888, targeting eight KAPs. Since 2010, tests for HIV, syphilis and hepatitis C virus (HCV) infections have been performed routinely for all participants recruited during the surveillance period [8].

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In recent years, harm reduction programmes have been implemented throughout China [9.10]. The two largest of these aimed at harm reduction among DUs are the national methadone maintenance treatment (MMT) programme and needle exchange programme (NEP). MMT was introduced in 1993 [11], and was backed as a treatment programme from 2004 [12], then expanded quickly from its original eight clinics in 2004 to 320 clinics in 2006 and to 738 clinics in 2011. Opiate users covered by the MMT programme increased cumulatively from 1209 in 2004 to 37 345 in 2006 and to 344 254 in 2011 [3]. NEP was first introduced to increase the commercial availability and accessibility of needles [13]. Until a successful intervention trial in Guangdong and Guangxi [14,15], 91 pilot NEP sites were established in 2003 [16]. Similar to the MMT programme, expansion of NEP also occurred after 2006, increasing to 775 sites in 2007 and to 937 sites in 2011. IDUs covered by NEP average approximately 50 000 monthly since 2007 [3,17,18].

Roughly 10 years have passed since the initiation of these harm reduction programmes in China. Although studies of the initial pilot MMT and NEP programmes provided sufficient evidence of their effectiveness to merit their expansion nation-wide [19,20], their impact on the national HIV epidemic has not yet been reported. In the present study, we collected information from China's national HIV/AIDS case reporting system and sentinel surveillance system. We aimed to describe the characteristics and trends of the HIV epidemic among the DU, specifically the IDU, population in China over time (1995–2011) and, secondarily, to describe changes in the behaviour of DUs and IDUs over the same time-period.

METHODS

For the purposes of this study, a DU was defined as a drug user who used 'traditional drugs' (i.e. heroin, opium, opiate analgesics, cocaine and marijuana, as defined in China's sentinel surveillance guidelines) [20–22] and/or the newer, so-called 'nightclub drugs' [i.e. methamphetamine, ketamine, 3,4-methylenedioxy-N-methylamphetamine (MDMA) and Magu pills (a mixture of methamphetamine and caffeine)] by any means, including via injection. An IDU was defined as a DU who had injected drugs in their life-time. A current IDU was defined as an IDU who had injected drugs in the past month. A non-IDU was defined as a DU who had never injected drugs in their life-time.

Data sources and collection

All data included in this study were collected from the national HIV/AIDS CRS and the national HIV sentinel

surveillance system. The data sets from each system were evaluated separately; they were not combined.

Issued in 1989, the Law of the People's Republic of China on the Prevention and Treatment of Infectious Diseases added HIV/AIDS to its listing of mandatorily reportable diseases. At hospitals, HIV testing became mandatory prior to surgery and other invasive diagnostic and treatment procedures. HIV testing was also conducted at voluntary counselling and testing sites, located typically at hospitals and local Center for Diseases Control and Prevention (CDC) clinics. Once HIV infection is confirmed [screening by the two-enzyme-linked immunosorbent assay (ELISA) method and a confirmatory test by Western blot], the provider informs the patient of their HIVpositive status and conducts a face-to-face interview to complete a case-reporting form (CRF). Name, identification number, age, gender, occupation and current address are collected as well as self-reported history of high-risk behaviour. The probable transmission route is judged by the provider based on the patient's behavioural history. Thus, HIV-positive patients who self-report ever injecting drugs in their life-time during this initial CRF interview are judged to be infected with HIV via drug injection. Completed CRFs are submitted to the local CDC. In 2005, the CRS was embedded into the newly launched, web-based China Information System for Disease Control and Prevention, and HIV/AIDS cases are required to be reported online within 24 hours of identification [4]. The National Center for AIDS/STD Control and Prevention (NCAIDS) became responsible for regular auditing of data timeliness and quality.

The national HIV sentinel surveillance system was established in 1995 for the purpose of estimating HIV prevalence among high-risk groups, including DUs [5]. From 2011, 303 sentinel sites have focused primarily on DUs. Each year, an anonymous cross-sectional survey is conducted from April to June. For DU surveillance, specifically, participants are recruited by a stratified snowball method within DU communities, regardless of their HIV testing history and serostatus. Seeds are selected from among peer educators, community volunteers and others with extensive networks among the DU community. DUs already enrolled in methadone maintenance treatment (MMT) are excluded from sampling unless they have had at least one positive urine opiate screen result in the 3 months prior to the survey [20]. Target sample size for each DU sentinel site is 400. A uniform questionnaire is used for face-to-face interviewing of participants at all DU sentinel sites. At each interview, participants provide demographic information, drugs used and drug use behavioural information (i.e. types of drugs used, method of drug use in the past month and frequency of drug use) and a blood sample for HIV testing (screening test by the two-ELISA method) [21]. Syphilis testing was added in

2004 [7] and HCV testing was added in 2009 [22]. Prior to 2007, an EpiData version 3.1 (EpiData Association, Odense, Denmark) database was used for data input and management and all sentinel surveillance data were reported to NCAIDS by encrypted EpiData files. In 2007, a client software program was designed for data input and management, which is uploaded in real time to the web server at NCAIDS.

Data from both the national HIV/AIDS CRS and the national HIV sentinel surveillance system were used for this study. From the CRS, all records from 31 December 2011 were downloaded, but only those cases judged to be infected by injecting drugs were selected (i.e. this data set included IDUs only) and sorted by the year in which the case was first reported. From the sentinel surveillance system, all annual data from 31 December 2011 were downloaded for all DUs (i.e. this data set included all DUs, meaning IDUs and non-IDUs together). Those DU sentinel sites having sample sizes of fewer than 50 respondents in a single year were excluded from analysis.

Data analysis

Annually reported HIV/AIDS cases from the national HIV/AIDS CRS were sorted by county and displayed on maps colour-coded for numbers of cases using ArcInfo software (ArcMap[™] version 10.0; ESRI®, Redlands, CA, USA) to examine geographic changes in the HIV epidemic in China over time (Fig. 1).

HIV prevalence data came from the national sentinel surveillance system, specifically DU sentinel sites, which include data for all DUs (both IDU and non-IDU). HIV prevalence estimates from the national HIV sentinel surveillance system were calculated for the five most affected provinces and for all other provinces, separated into IDU and non-IDU groups (Fig. 2). Prevalence was calculated as the number of HIV-positive respondents as the numerator and the total number of respondents as the denominator and presented with 95% confidence intervals (CIs).

HIV prevalence among IDUs alone, nation-wide, is taken from annual sentinel surveillance data. To illustrate the possible impact of the expansion of the number of DU sentinel surveillance sites in China after

Figure I Geographic distribution of the number of HIV/AIDS cases reported among injecting drug users (IDUs) in 1995, 2000, 2005 and 2011. Variation in colour indicates the numbers of reported HIV/ AIDS cases by county. Lines within the map indicate province and county borders. All data presented here originated from the national HIV/AIDS case reporting system (CRS) and include those cases judged to be infected with HIV via unsafe injecting behaviours, thus all cases included are defined as IDUs





Figure 2 Graphic representation of changes in HIV prevalence over time (1998–2011) among injecting drug users (IDUs) [drug users (DUs) judged to have become infected with HIV via unsafe injecting practices] and among non-IDUs (DUs judged to have become infected with HIV via transmission routes other than injecting drugs) in the five most affected provinces (red)—Yunnan, Guangxi, Sichuan, Xinjiang and Guangdong—compared to all other provinces (green). Bars at each data point represent the 95% confidence intervals of prevalence. All data presented here were obtained from the national HIV sentinel surveillance system



Figure 3 Graphic representation of the trends in HIV prevalence among injecting drug users (IDUs) only: nation-wide (all active sentinel sites; red), and at the seven sites having longest surveillance time-span form 1998 to 2011 (purple). Bars at each data point represent the 95% confidence intervals. The numbers of active sentinel sites for each year appear in the form of light green bars to highlight the rapid scale-up of China's national sentinel surveillance system after 2005. Those drug user (DU) sentinel sites having sample sizes of fewer than 50 respondents in a single year were excluded from analysis (target sample size is 400 respondents per sentinel site). All data presented here came from the national HIV sentinel surveillance system

2005, the overall HIV prevalence among IDUs only for all active sites each year and for the seven sites with the longest history (data collected at these seven sites each year between 1998 and 2011) was plotted, along with the total number of active sentinel sites each year (Fig. 3).

Data related to drug use behaviours among all DUs come from national sentinel surveillance data collected after 2004. To calculate the proportions of 'nightclub drug' users, all DU respondents in the survey year was the denominator and all DU respondents who self-reported using 'nightclub drugs' was the numerator. Similar calculations were conducted for each individual specific 'nightclub drug' (Fig. 4). To calculate the proportions of 'traditional drug' users, all IDU respondents in the survey year was the denominator and all IDU respondents who self-reported using 'traditional drugs' was the numerator. To calculate the proportions of current IDUs who engaged in recent needle sharing, all current IDU respondents in the survey year was the denominator and all IDU respondents who self-reported needle sharing in the past month was the numerator (Fig. 5).



Figure 4 Graphic representation of the rapid increase in the prevalence of 'nightclub drug' users among drug user (DU) sentinel surveillance survey respondents from 2004 to 2011. Prevalence of use of the four most popular 'nightclub drugs' among 'nightclub drug' users. All data presented here came from the sentinel surveillance system. These data illustrate the very recent and rapid escalation in use of 'nightclub drugs' among 'nightclub drugs' and the diversity of drugs used. Due to the large sample size, 95% confidence intervals are not shown in the figure



Figure 5 Graphic representation of changes over time (2004–11) in the proportion of current injecting drug users (IDUs) (i.e. IDUs who self-reported injecting drugs in the most recent month prior to the sentinel surveillance survey) among 'traditional drug' users and the proportion of current IDUs who self-reported recent needle-sharing behaviour. The scaling-up of methadone maintenance treatment (MMT) and needle exchange programme (NEP) harm reduction programmes is also shown. All data were obtained from the national HIV sentinel surveillance system. Needle-sharing information was not collected before 2006. Due to the large sample size, 95% confidence intervals are not shown in the figure

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Pearson's correlation tests were used to investigate a possible relationship between trends in DU behavioural characteristics and the expansion of harm reduction programmes. All data analysis was performed using with SPSS software (version 18.0.0 SPSS, Inc., Chicago, IL, USA).

Ethical approval

This study was reviewed and approved by the Institutional Review Board of NCAIDS, China CDC. Data used in this study were collected via routine monitoring and surveillance. All personal identifiers were removed from the final national HIV/AIDS CRS data set. National sentinel surveillance data were collected anonymously and were thus already free of name-linked information. Therefore, written informed consent was not required.

RESULTS

Geographic trends in HIV prevalence

Figure 1 shows maps of the distribution of identified HIV/ AIDS cases among IDUs. In 1995, the epidemic was highly concentrated in southwestern Yunnan province. Five years later, in 2000, the HIV epidemic among IDU spread to neighbouring southwest provinces and another subepidemic began in Xinjiang Uygur Autonomous Region in the northwest. In 2005, the epidemic among IDUs spread to the northeast, and by 2011 HIV-positive IDU had been reported in 1703 prefectures nation-wide. After two decades of expansion of the HIV epidemic among IDUs in China, it is clear that southwest provinces (Yunnan, Sichuan, Guangxi and Guangdong) and Xinjiang Uygur Autonomous Region in the far northwest still bear a disproportionately large HIV burden relative to the rest of the country.

This geographic pattern in the distribution of the HIV epidemic is also reflected in data for all DUs (IDUs and non-IDUs) from the national sentinel surveillance system. As shown in Fig. 2, the prevalence of HIV among both IDUs and non-IDUs was much higher and peaked earlier in the five most affected provinces than in all other provinces combined. HIV prevalence among IDUs in the five provinces peaked in 1999 at 30.3% (95% CI = 28.6, 32.1) and decreased to 10.9% (95% CI = 10.6, 11.2) by 2011. By contrast, HIV prevalence among IDUs in all other provinces peaked in 2003 at 5.2% (95% CI = 4.7, 5.7) and then declined to 1.6% (5% CI = 1.4, 1.7) by 2011. Non-IDUs in the five most affected provinces also had much higher HIV prevalence compared to all other provinces from 1998 to 2011. HIV prevalence among non-IDU in five provinces reached 6.3% (95% CI = 5.5, 7.1) in 2005 and then declined to 2.2% (95% CI = 2.0,

2.4) by 2011, whereas HIV prevalence in all other provinces was less than 1% over the entire time-period.

Nation-wide trends in HIV prevalence

Figure 3 shows the annual nation-wide changes in HIV prevalence among IDUs from 1998 to 2011. After a rapid increase before 2000, HIV prevalence among IDUs nation-wide increased moderately to a peak of 10.9% (95% CI = 10.4, 11.4) in 2004. Since 2004 there has been a decline, and in 2011 nation-wide HIV prevalence among IDUs was 6.4% (95% CI = 6.2, 6.6). This period of lower prevalence from 2006 to 2011 coincided with the rapid scale-up of sentinel surveillance sites, and thus could have resulted merely from a dilution effect (i.e. sentinel sites were opened in high-prevalence areas first, and low-prevalence areas later). To eliminate the potential dilution effect, data from the seven sentinel sites with the longest history (having HIV prevalence survey data each year from 1998 to 2011) were overlaid on the same graph. These data show that HIV prevalence at these seven sites reflects a trend similar to the national data, as HIV prevalence for these seven sites declined from 6.3% (95% CI = 5.0, 7.6) in 2003 to 1.6% (95% CI = 1.0, 2.2) in 2011.

Nation-wide trends in drug use behaviours

As shown in Fig. 4, DUs engaging in 'nightclub drug' use increased from 1.3% in 2004 to 24.4% in 2011. The most popular of these drugs were ketamine and methamphetamine, which together accounted for the drugs of choice among the majority of 'nightclub drug' users from 2004 to 2011.

Trends in injecting behaviour in the past month among 'traditional drug' users and needle sharing among current IDUs (injecting in the past month) are shown in Fig. 5. While the proportion of IDUs among 'traditional drug' users decreased slightly from 47.2% (95% CI = 47.1, 47.3) in 2004 to 44.8% (95% CI = 44.9, 44.9) in 2011, the proportion of current IDUs who reported sharing needles in the past month decreased from 19.5% (95% CI = 19.4, 19.6) in 2006 to 11.3% (95% CI = 11.2, 11.4) in 2011. Pearson's correlation test results show that this decrease in needle sharing was correlated with the expansion of MMT ($r_{(4)} = -0.94$, P = 0.003, data not shown).

Although MMT participants are excluded from sentinel surveys unless they have at least one positive urine opiate screen result in the 3 months prior to the survey, we compared the injection behaviour among sentinel survey respondents who had ever participated in MMT to respondents who had never participated in MMT. We found that among DUs who had participated in MMT, fewer were current IDUs (74.3%, 95% CI = 74.1, 74.5) compared to DUs who had never participated (78.3%, 95% CI = 78.1, 78.5, χ^2 = 133.15, *P* < 0.001, data not shown).

DISCUSSION

We found that despite the broad geographic expansion of China's HIV epidemic during the past two decades, most cases are still concentrated in the five provinces initially affected by the epidemic—Yunnan, Sichuan, Guangxi, Guangdong and Xinjiang Uygur Autonomous Region. This geographic pattern has also been observed in several previous studies [23,24] and mirrors global drug trafficking activities, as the majority of heroin and opium in China was brought in from Myanmar into Yunnan, or from Vietnam into Guangxi, and a small proportion from the Middle East into Xinjiang [25]. The easy accessibility of drugs in these five provinces gave rise to the rapidly expanding HIV epidemic among IDUs in these areas.

As the epidemic matured over time, HIV also spread to non-IDUs, probably via commercial and non-commercial sex. This is evidenced by Yunnan, Guangxi and Xinjiang provinces reporting a high HIV prevalence among female sex workers and their clients [26,27], as well as pregnant women [28,29], as early as 2004. Similar secondary sexual transmission patterns have been observed in several other countries in the region. For example, India and Pakistan have reported secondary HIV transmission among non-IDU spouses of HIV-positive IDU [30,31], and a molecular epidemiological study has identified linkages among the sexual and injecting drug use epidemics in Thailand [2]. Efforts to control the spread of HIV within and outside these most-affected five provinces must therefore be focused upon both IDUs and their sex partners.

We also found that changes in drug use-related behaviours in recent years were correlated with the rapid scale-up of harm reduction programmes in China from 2006 to 2011. For example, needle sharing behaviour among current IDUs decreased significantly over this period, an effect which has a significant correlation with the expansion in coverage of MMT. Our results, which associate the positive influence of harm reduction programmes with decreased dangerous drug use-related behaviours, are supported by past research, including several small studies which describe the successes of MMT and NEP as harm reduction interventions [19,32]. Larger studies have variously reported a significant decrease in both high-risk sexual and drug use behaviour following MMT enrolment in Yunnan [14,15,33,34].

DUs already enrolled in MMT are excluded from sentinel surveys unless they have at least one positive urine opiate screen result in the 3 months prior to the survey. This sampling criterion limited our analysis on the effect of MMT on DU behaviours. However, we found that a smaller proportion of DUs who had ever received MMT services were current IDUs, compared to DUs who had never received MMT services. Some international studies have also found similar trends and changes, where MMT and NEP not only reduced drug use-related behaviours significantly, but also decreased HIV prevalence in Vietnam, Iran and other countries [35–37]. In addition to the rapid scale-up of harm reduction programmes in China, the Chinese government has also adopted more effective policies and cooperative measures targeting drug abuse problems, such as the Cross-Border Project, which was implemented in both northern Vietnam and southern China [38]. Legislation concerning compulsory rehabilitation centres has also changed since 2011, which may influence the drug use-related behaviours of drug users.

In order to prolong the life expectancy of AIDS patients living in rural regions of China, the Chinese government initiated the national free antiretroviral therapy (ART) programme in 2002 [39]. More than 80 000 patients had received ART from 2002 to the end of 2009. and the free ART programme had already expanded to 28 provinces of China by the end of 2005 [39,40]. Earlier initiation of ART and increased treatment coverage can reduce the mortality rate of HIV-infected people significantly, but IDUs in China were less likely to receive ART compared with those infected with HIV through sexual behaviour and plasma donation [41]. Although some studies indicate that ART may contribute to a reduction in risk behaviours associated with HIV transmission [42]. and help HIV-infected people to perceive the importance of safer practices [43], studies indicating a valid relationship between the free ART programme and change in risk behaviours among DUs in China could not be found. None the less, the combined influence of MMT, NEP, ART and other systematic interventions on reducing risk behaviours among DUs should not be ignored.

Finally, we found a decreasing trend in HIV prevalence among DUs. Although the rapid scale-up of sentinel sites over the same time-period and the limitations inherent in non-probability sampling methods reduced the representativeness of our finding, we also examined data from the seven sentinel sites that existed and actively collected annual survey data in each consecutive year during the time-period we investigated. These data confirmed the decreasing trend of HIV prevalence among DUs. The observed decline in DU HIV prevalence may be due, in part, to a decline in the population of infected DUs and a low rate of new infections among this population [34]. Although the coverage of China's national ART programme expanded greatly after 2003, the high degree of late diagnosis of HIV infection, co-infection with multiple viral strains, serious comorbidities such as HCV infection and increasing drug resistance gave rise to increasing numbers of AIDS patients [44] and disproportionately

high HIV/AIDS-related mortality among DUs [45]. Government epidemic estimation efforts reported that new infections among DUs were fewer than 8000, yet deaths among DUs were more than 9000 in 2011 [3]. Another potential reason for the decrease in HIV prevalence may be refusal to participate in the sentinel surveys and/or avoidance of testing among DUs. A recent study reported a difference in HIV prevalence among DUs responding to sentinel surveillance surveys and DUs in the broader community [46].

Our study has some limitations. As an observational study, we describe trends and identify correlations, but are unable to examine causal relationships. Furthermore, sentinel surveillance data are based on a stratified snowball sampling method and thus may not be representative of the broader DU population in China.

In summary, while the HIV epidemic among DUs in China has expanded during the past two decades, it is still highly concentrated in five provinces. HIV prevalence and high-risk behaviour among IDU have declined and might be correlated with the scale-up of national harm reduction efforts. However, the recent rapid increase in the use of 'nightclub drugs' presents a new challenge for China's Ministries of Public Security and Health in their joint effort to curb both drug use and the HIV epidemic.

Declaration of interests

None.

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