

Use of Population-Based Surveys for Estimating the Population Size of Persons Who Inject Drugs in the United States

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Background. In the United States, injection is an increasingly common route of administration for opioids and other substances. Estimates of the number of persons who inject drugs (PWID) are needed for monitoring risk-specific infectious disease rates and health services coverage.

Methods. We reviewed design and instruments for 4 national household surveys, 2012–2016, for their ability to produce unbiased injection drug use (IDU) prevalence estimates. We explored potential analytic adjustments for reducing biases through use of external data on (1) arrest, (2) narcotic overdose mortality, and (3) biomarker-based sensitivity of self-reported illicit drug use.

Results. Estimated national past 12 months IDU prevalence ranged from 0.24% to 0.59% across surveys. All surveys excluded unstably housed and incarcerated persons, and estimates were based on <60 respondents reporting IDU behavior in 3 surveys. No surveys asked participants about nonmedical injection of prescription drugs. Analytic adjustments did not appreciably change IDU prevalence estimates due to suboptimal specificity of data points.

Conclusions. PWID population size estimates in the United States are based on small numbers and are likely biased by undercoverage of key populations and self-report. Novel methods as discussed in this article may improve our understanding of PWID population size and their health needs.

Keywords. PWID; injection behavior; population-based surveys

In the United States, injection is an increasingly common and highrisk route of administration for prescription and illicit opioids [1– 5] as well as other drugs such as methamphetamine [6, 7]. Unsafe injection drug use (IDU) behaviors increase risk for bloodborne infectious diseases such as hepatitis C virus (HCV) and human immunodeficiency virus (HIV), making these infectious diseases secondary but deleterious consequences of the opioid epidemic for persons who inject drugs (PWID) [8, 9]. Due to the stigmatized and illicit nature of nonmedical IDU, population-level prevalence is difficult to measure using survey methods typically used to monitor health-related behaviors. However, understanding the national and local scopes of IDU behavior is critical for informing infectious disease prevention efforts among PWID and to help ensure the success of ambitious strategies to eliminate HIV and HCV in the United States [10–13].

Information about the size and characteristics of the PWID population is needed to inform strategies to reduce

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IDU-associated infectious disease risk at multiple phases of intervention design and monitoring. First, PWID population size estimates are needed to enable data-informed resource allocation and program planning decisions. Second, as prevention and treatment interventions are implemented, estimates of PWID population size can help monitor progress toward coverage goals. Last, PWID population size is needed to monitor infectious disease rates among the at-risk population most in need of health interventions [14, 15].

Evidence of increasing IDU prevalence amplifies the need for current, accurate PWID population size estimates. The national drug overdose death rate increased from 6.1 to 21.7/100 000 during 1999–2017 [16]. Acute HCV infections increased by more than 3.5-fold during 2010–2017, mostly among young adults [17], and IDU-associated increases in HIV infections have recently been observed in rural areas where HIV is rare [1, 2, 18–20]. National increases in infectious conditions typically associated with IDU behavior, such as infectious endocarditis and central nervous system abscesses, have been observed in emergency room and hospital discharge data [21–24]. Indirect but highly correlated indications of IDU behavior suggest further increases in IDU-associated infections are likely.

In the United States, like many health-related behaviors, IDU is measured and monitored primarily using household surveys,

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and prevalence estimates are derived from data from 1 or more surveys [25]. Estimates derived from survey data are generally subject to several types of bias (Figure 1) [26], which may variably affect estimates depending on the condition of interest. Two sources of bias threaten the ability of the study sample to reflect the underlying population of interest: (1) selection bias: when a sampling frame is suboptimally representative of the population, as is the case when, for example, homeless populations who may have relatively high IDU prevalence are not captured in a household survey; and (2) nonresponse bias: when nonresponse of sampled persons is differential by the condition of interest [27, 28]. Two levels of information bias threaten the ability of collected data to represent the construct of interest: (1) questions that are suboptimally designed to measure the construct of interest in the intended sample; and (2) inaccurate participant responses due to misunderstanding of survey questions, limited time or ability to complete the survey, or a desire to provide responses that are socially acceptable or preferred by the interviewer. Due to stigma and legal implications surrounding IDU behavior, response bias is particularly problematic [27, 28]. Random error may further distort estimates, particularly in the context of small sample sizes.

To assess the comparative utility of household survey data sources for estimating PWID population size, we qualitatively evaluated how these biases may influence IDU prevalence estimates from national surveys currently collecting such data. We then conducted a feasibility assessment of potential adjustments to survey-derived estimates using established analytic methods and data external to household surveys. All proposed adjustments have either been previously used in other contexts or are extensions to methods previously used. We conclude with a discussion of possible improvements to survey estimates as well as emerging methods that may be used to estimate PWID population size.

METHODS

We examined 4 national population-based surveys in terms of ability to produce complete, unbiased, nationally representative measures of current nonmedical IDU prevalence. All 4 were US Department of Health and Human Services (HHS) surveys: the National Survey on Drug Use and Health (NSDUH, years 2015–2016) [29], the National Health and Nutrition Examination Survey (NHANES, 2013–2016) [30, 31], the National Survey of Family Growth (NSFG, 2013–2015) [32], and the National Institutes of Alcohol Abuse and Alcoholism (NIAAA)-supported Epidemiologic Survey on Alcohol and Related Conditions-III (NESARC-III, 2012–2013) [33]. Data from NSDUH, NHANES, and NSFG are in the public domain, and NESARC-III data were obtained through a data usage agreement with NIAAA for use of a limited access dataset.

For each survey, we abstracted and compared sampling frames and over-sampling strategies, interview modalities,

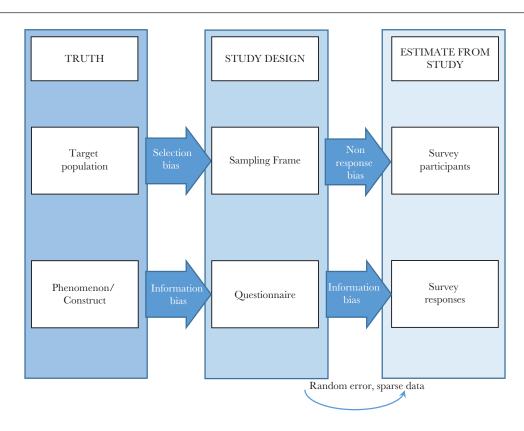


Figure 1. Framework for evaluating bias in estimates from survey research. Adapted from Hulley et al, Designing Clinical Research, 4th ed [26].

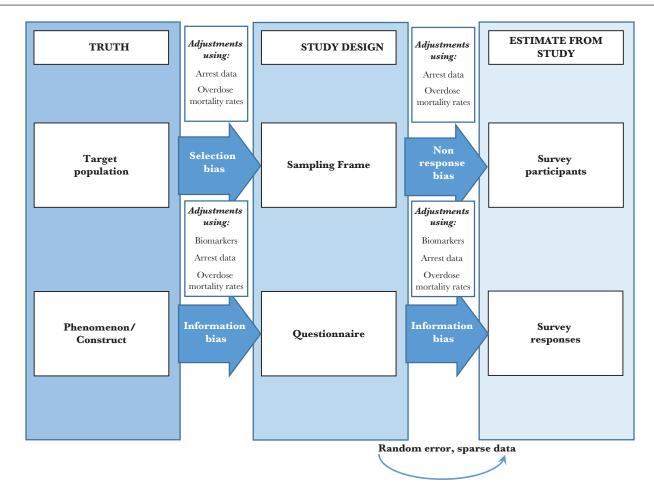
questionnaire item(s) language, and relevant skip patterns. In order to gauge to what extent random error due to small sample sizes and influential data points may affect estimates and to assess potential for producing stratified prevalence estimates, we compared unweighted numbers of persons reporting past year IDU behavior across surveys. We computed weighted prevalence of past year IDU prevalence with associated 95% confidence intervals, limiting estimates to persons aged 18–64 years with the exception of the NSFG-derived estimate, due to an upper age limit of 44 years in the survey design.

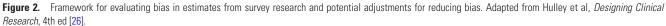
Next, using NSDUH data, we explored the feasibility of making 3 adjustments to population-based estimates of current IDU behavior in order to reduce biases identified (Figure 2). Where possible, we explored use of these adjustments within sex, age, and race/ethnicity strata. Stability of each stratified proportion, *p*, was assessed using NSDUH guidelines, relative standard error $[-\ln(p)] < 0.175$ [34].

Biomarker-Based Adjustment Method

To address information biases and threats to internal validity of self-reported IDU (p_{IDU}), we assessed feasibility of computing a standard multiplier for sensitivity of self-reported use of illicit drugs for use in adjusting misclassification of selfreported IDUp within demographic strata [14, 35]. We conceptualized such a multiplier as $\hat{p}_{corrected} = \hat{p}_{IDU} \times \frac{1}{sensitivity_{SR}}$, where *sensitivity*_{SR} is the number of individuals reporting such drug use within a specific recall period, divided by the total number testing positive for biomarkers of drug use.

We conducted a literature review to identify published estimates of the sensitivity of self-reported illicit (nonmarijuana) drug use. We searched studies published during 1995–2019 and indexed using PubMed. Search terms used were: [selfreport(ed), report(ed), sensitivity, accuracy, veracity, validity], [drug (use)], [illegal, illicit, injection], [urinalysis, biomarker, bioassay, screen]. The primary inclusion criterion was reported sensitivity of self-reported nonmarijuana drug use. We excluded studies in drug treatment or rehabilitation settings and those conducted outside the United States. For studies meeting inclusion criteria, we abstracted relevant data and qualitatively evaluated whether a standard multiplier could be computed for improving IDU prevalence estimates.





Arrest Data Adjustment Method

Second, to address selection, nonresponse, and information biases in prevalence estimates, we explored possible use of a multiplier adjustment previously proposed by the National Institute on Drug Abuse. This method uses self-reported arrest and IDU in conjunction with an external estimate of the number of persons arrested in the United States during the past year [36]. Using NSDUH data, we estimated weighted prevalences of IDU in the past year among persons who did $(\hat{p}_{IDU|arrest})$ and did not report being arrested $(\hat{p}_{IDU|\sim arrest})$ during the past year, overall and stratified by sex and age. Arrest data were not available by the same race/ethnicity categories used by NSDUH. We then scaled the prevalence of IDU to the US population by applying, within demographic strata, $\hat{p}_{IDU|arrest}$ and $\hat{p}_{IDU|\sim arrest}$ from NSDUH to the total persons in the United States aged 18-64 years who were (N_{arrest}) and were not $(N_{\sim arrest})$ arrested during the past year using external data from the 2016 Federal Bureau of Investigation Uniform Crime Report [37]. This multiplier is summarized by $\hat{p}_{corrected} = \frac{(\hat{p}_{IDU|arrest} \times N_{arrest}) + (\hat{p}_{IDU|\sim arrest} \times N_{\sim arrest})}{N_{arrest} + N_{\sim arrest}}.$

Because information was available only for total number of arrests, rather than number of persons arrested, we followed previous guidance on application of this method and divided the total number of arrests among persons aged 18–64 years in 2016 by the average number of arrests reported per person in NSDUH within sex and age strata [36]. The number of persons not arrested in 2016 was calculated by subtracting the number of persons arrested from the total population in 2016 [38]. Stratum-specific products of $\hat{P}_{corrected} \times$ population totals (ie, $N_{arrest} + N_{\sim arrest}$) were summed for an adjusted PWID population size.

Narcotic Overdose Mortality Adjustment Method

Last, to address selection, nonresponse, and information biases in IDU prevalence estimates, we used temporal growth in narcotic overdose mortality rates as an external data multiplier. Within sex, age, and race/ethnicity strata, we applied the ratio of overdose death rates among US adults aged 18–64 years during 2015–2016 ($\mu_{2015-2016}$) to that during 2010–2011 ($\mu_{2010-2011}$) to the NSDUH IDU prevalence estimate in 2010–2011 (\hat{P}_{IDU} , 2010–2011). This multiplier is summarized as: \hat{P}_{IDU} , 2015–2016 = \hat{P}_{IDU} , 2010–2011 × ($\frac{\mu_{2015-2016}}{\mu_{2010-2011}}$).

Narcotic overdose death rates were estimated separately for 2010–2011 and 2015–2016 using National Vital Statistics System Multiple Cause of Death Mortality data and US Census Intercensal data. Multiple Cause of Death Mortality Microdata files (2010–2011 and 2015–2016) were obtained from the National Vital Statistics System. These data include individual death records for persons aged 18–64 years who lived in a US state or the District of Columbia and contained International Classification of Diseases, Tenth Revision (ICD-10) codes for multiple underlying causes of deaths. Narcotic overdose death was defined using the ICD-10 codes for unintentional poisoning by and exposure to narcotics and psychodysleptics (hallucinogens) (X42), unknown intention poisoning by and exposure to narcotics and psychodysleptics (hallucinogens) (Y12), unintentional poisoning by and exposure to other and unspecified drugs, medicaments, and biological substances (X44), or unknown intention poisoning by and exposure to other and unspecified drugs, medicaments, and biological substances (Y14). This definition has previously been shown to be more specific to injection-related overdose deaths than alternatives [39].

RESULTS

The estimated overall past year prevalence of IDU among the civilian, noninstitutionalized US population aged 18–64 years ranged from 0.24% in NESARC (2012–2013) to 0.59% in NSFG (2013–2015, among 18–44 year olds) (Table 1). All surveys sampled only persons within households from the civilian, noninstitutionalized population. Unweighted numbers of persons reporting IDU in the past year varied greatly from 38 (NHANES) to 380 persons (NSDUH).

With regards to survey administration, all were administered using audio-computer assisted self-interview (ACASI) with the exception of NESARC. Questions used to capture past year injection and associated skip patterns varied greatly. NESARC and NSDUH asked participants whether and during what time frame they last injected if they previously reported use of prespecified drugs. NSFG asked everyone if they had injected in the past year, and NHANES asked about time since last injection among everyone who reported ever injecting. Questions from NSDUH, NSFG, and NHANES included language indicating the drugs in question were drugs not prescribed by a doctor or not specifically prescribed to the respondent.

NSDUH was the only survey in which an adequate number of respondents reported past year IDU behavior for computation of stratified prevalence estimates (n = 380), likely due to oversampling of persons aged 18–39 years. Other key advantages of the NSDUH survey are the use of ACASI and its focus on substance use behaviors, which requires specialized staff training for data collection among substance-using populations. For these reasons, NSDUH IDU prevalence estimates were used for feasibility assessment of the following correction factors.

Biomarker-Based Adjustment Method

Self-reported sensitivity of illicit, nonmarijuana drug use varied widely across published studies (Table 2) [40–46]. Biomarkers are used to validate drug use rather than mode of administration, so no studies evaluated sensitivity of self-reported injection behaviors. Seven studies met inclusion criteria; many were excluded because they were conducted in drug treatment or rehabilitation settings. Time frames for data collection ranged from 1996 to 2017.

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Survey	Year	Most Recent Weighted Estimate Among 18–64 Year Olds, ^a % (95% Cl)	Unweighted Number With Outcome	Question(s) Used	Population Frame	Interview Modality Populations Over (eg, Sampled ACASI)	v V Skip Pattern Considerations
National Epidemiologic Survey on Alcohol and Related Conditions (NESARC)-III	2012-2013	0.24 (. 1631)	8	Did you take ANY medicines or drugs that we just talked about (sedatives or tranquilizers, painkillers, cocaine or crack, stimulants, club drugs, hallucinogens, heroin, other) on your own by injection with a needle in the last 12 months?	Civilian, noninstitutionalized persons aged 18 y and older	Census blocks with CAPI more than 26% of the population who identify as Hispanic, Black, or Asian	Respondents were asked about injection only if they reported any drug use (of sedatives or tranquilizers, painkillers, cocaine or crack, stimulants, club drugs, hallucinogens, heroin, or other) apart from marijuana and/or inhalants/solvents
National Survey on Drug Use and Health (NSDUH)	2015-2016	0.41 (.35– .47)	380	How long has it been since you last used a needle to inject (cocaine, heroin, methamphetamine, stimulants, any other drug not prescribed to you)? (options: within the past 30 days, more than 30 days ago but within the past 12 months, and more than 12 months ago).	Civilian, noninstitutionalized persons aged 12 y and older	Persons aged 12–25 yACASI	Respondents were asked about past year injection only if they reported lifetime use of predetermined drugs (for 2012–2013, drugs were cocaine, heroin, metham- phetamine, or stimulants; for 2015–2016, drugs were cocaine, heroin, methamphet- amine, or any other drug not prescribed to them) using a needle
National Health and Nutrition Examination Survey (NHANES)	2013-2016	0.45 (.27– .63)	ő	How long ago has it been since you last used a needle to inject a drug not prescribed by a doctor? (Time periods of 12 months or less.)	Civilian, noninstitutionalized persons aged 12–69 y	Persons who identify ACASI as Hispanic, non- Hispanic Black, non-Hispanic Asians, non- Hapanic White, or Other persons at or below 130% fed- eral poverty level, and persons aged 80 y and over	Respondents were only asked if they indi- cated that they had ever used a needle to inject an illegal drug
National Survey of Family Growth (NSFG)	2013-2015	0.59 (.35– .82)	53	During the last 12 months, how often have you shot up or injected drugs other than those prescribed for you? By shooting up, we mean anytime you might have used drugs with a needle, by mainlining, skin-popping, or muscling.	Civilian ^b , noninstitutionalized persons aged 15–44 y	Persons who identify ACASI as Black or His- panic, or who are aged 15–19 y	Applicable for all respondents
Abbreviations: ACASI, audio-comp ^a Among 18-44 year olds in NSFG. ^b Includes military living off base.	sl, audio-comput olds in NSFG. ing off base.	Abbreviations: ACASI, audio-computer assisted self-interview; CAPI, computerassisted Among 18-44 year olds in NSFG. [*] Includes military living off base.	w; CAPI, compute	rassisted personal interviewing; Cl, confidence interval.	terval.		

Table 1. National Estimates of Injection Drug Use in Past 12 Months From Population-Based Surveys and Survey Characteristics

Study Year Methampher. Methampher. Harom 2010-2012 454 black MSM aged 18-40 y from Atlanta area 10 (22)* 0.54 (1975) 0.05 (1975) No Ledgerwood 1996-1997 839 midele-aged men from a cohort of individuals 0.44* 0.46* 0.78* 0.78* Ledgerwood 1996-1997 839 midele-aged men from a cohort of individuals 0.44* 0.46* 0.78* 0.78* Ledgerwood 1996-1997 839 midele-aged men from a cohort of individuals 0.44* 0.46* 0.78* 0.78* Ledgerwood 1996-1997 839 midele-aged men from a cohort of individuals 0.44* 0.46* 0.78* 0.78* Ledgerwood 1996-197 839 midele-aged men from a cohort of individuals 0.44* 0.46* 0.78* 0.78* Leddich et al 2001-2002 1996-138* 9.8* aged 31-40 mine, only 2 0.14* 0.46* 0.79* 0.66 Leddich et al 2001-2002 101* 101* 101* 0.14* 0.25* 11* 0.14* 0.25*					
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1936–1937 B33 middle-aged men from a cohort of individuals diaployed to Vietnam and a matched nonveteran group, 16.3% black, 78.2% white 0.44° 0.46° 2001–2002 627 Chicago residents aged 18–40 y selected by multistage area probability design. 42.8% aged 18–25 y; 25.5% aged 26–30 y; 31.8% aged 31–40 y; 36.6% black; 33.2% white; 22.1% Hispanic; 9.2% other NA (tested for reside inc, only 2 Unime: 0.46° Unime: 0.46° Virine: 0.36° 2015–2017 1029 individuals aged 16–29 y, assigned male at birth. 0.37° from the Chicago. IL area who identify as agy or bisexual and/or reported sex with a man in the last 12 mo; 33.1% black; 24.9% white, 29.8% 0.34° 0.37° 0.34° 2015–2017 1029 individuals aged 16–29 y, assigned male at birth. 0.37° 0.36° 0.37° 0.36° 2015–2017 1029 individuals aged 16–29 y, assigned male at birth. 0.37° 0.36° 0.36° 0.56° 2015–2017 1029 individuals aged 06–29 y, assigned male at birth. 0.37° 0.36° 0.56° 0.56° 2015–2017 1029 individuals aged 16–29 y, assigned male at birth. 0.37° 0.36° 0.56° 0.56° 2015 83 pregnant women from New Mexico; 80% Latins. 0.37° 0.36° 0.56° 0.56° 7% American Indian; 13% other 0.37° 0.37° 0.37° 0.36° 0.37° 0.36°	0.82 (9/11) ^a 0.84 (16/19)	م(0/0)		Past 12 mo	
2001-2002 627 Chicago residents aged 18-40 y selected by multistage area probability design. 42.8% aged 18-25 y, 25.5% aged 26-30 y; 31.8% aged 31-40 NAI (tested for Hair: 0.24° ampheta- 18-25 y, 25.5% aged 26-30 y; 31.8% aged 31-40 18-25 y, 25.5% aged 26-30 y; 31.8% aged 31-40 mine, only 2 Urine: 0.46° y; 35.6% black; 33.2% white; 22.1% Hispanic; 9.2% other Value 2% other 9.2% other Fasted pos- for MDMAJ Lorine: 0.30° Urine: 0.30° Urine: 0.30° 2015-2017 1029 individuals aged 16-29 y, assigned male at hirth 0.33° from the Chicago. IL area who identify as gay or bisexual and/or reported eas white, 29.8% white, 29.8% 0.37° Urine: 0.30° 2015-2017 1029 individuals aged 16-29 y, assigned male at hirth 0.33° 0.37° Urine: 0.30° 2015-2017 1029 individuals aged 16-29 y, assigned male at hirth 0.33° 0.37° Urine: 0.30° 2015-2017 1029 individuals aged 16-29 y, assigned male at hirth 0.33° 0.37° Urine: 0.30° 2015-2017 1029 individuals aged 16-29 y, assigned male at hirth 0.33° 0.37° Urine: 0.30° 2015-2017 1029 individuals aged 16-29 y, assigned male at hirth 0.33° 0.37° Urine: 0.30° 2015-2017 1029 individuals aged 16-29 y, assigned male at hirth 0.33° 0.37° Urine: 0.30° 2015 1050 individuals 0.53° 0.53° 2016 0.37° 0.37°	0.44°	8 ^{6,6} Hair	Cocaine, cocaine, benzoyleogonine, or cocaertrylene Opiates: codeine, morphine glucuronide, or 6-acetylmorphine Methamphetamine: ampheta- mine and methamphetamine Radioimmunoassay, GC-MS. 5 ng drug/10 g hair cutoff for GC-MS. testino	Past 90 d	50% of veterans screened positive for drugs at enrol- ment in 1971°
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2012 B3 pregnant women from New Mexico, 80% Latina; 0.37 ^{a,c} 0.33 ^c 7% American Indian; 13% other 0.29 ^c 0.29 ^c	0.37° 0. 0.53° 0. 0.50° 0.	0.27 ^{b.c} Urine 0.27 ^{b.c} 0.727 ^{b.c}	Methamphetamine: D-methamphetamine (500 ng/L) Cocaine: benzoylegonine (150 ng/L) Heroin: morphine (300 ng/L)	Window period lower limit methamphetamine: 3 d, cocaine: 1 d, opiates: 1 d Window period upper limit methamphetamine: 3 d, opiates: 4 d Past 6 mo Past 30 d	
8/18)	0.37 ^{b.c} 0.33 ^c 0.29 ^c 0.44 (8/18) 0.44 (21/48)	0.478 ^{b.c} Urine 0.17 ^{b.c} Saliva	Not specified Not specified	Occasional use (once per month or less) Regular use (more frequently than once per month) Past 72 h	Women were selected on his- tory of substance abuse ^o
al [42] 1999 29	0.67 (42/89) tly	Urine	Not specified	Past 30 d Past 7 d	Drugs tested for were not specified ^c

Table 2. Sensitivity of Self-Reported Illicit Drug Use Estimates From Research Studies in Nondrug-Treatment Settings

The studies were heterogeneous across study populations, biomarker(s) used (urine, hair, or saliva), drugs tested for, assays used for testing, and detectable time window of drug use due to recall periods for survey questions, biomarkers, and assays used. Generally, the numbers of people reporting nonmarijuana drug use was small. Self-report sensitivity estimates for methamphetamine ranged from 0.28 to 1.00, estimates for cocaine ranged from 0.14 to 0.60, and estimates for heroin ranged from 0.17 to 0.78. Due to heterogeneity in study designs and associated sensitivity estimates and the inability to stratify sensitivity estimates by demographic characteristics, we concluded it was imprudent to compute a combined correction factor for adjusting household survey IDU estimates.

Arrest Data Adjustment Method

Self-reported IDU prevalence was considerably higher among persons reporting arrest during the past year than among those not reporting arrest (5.55% vs 0.27% overall) (Table 3). Overall, we estimated 9.1 million people, or 4.5% of persons aged 18–64 years, were arrested during 2016 [37]. Applying the percentage of persons self-reporting IDU among arrested and nonarrested persons and summing the products across strata of arrested and nonarrested populations resulted in an adjusted overall IDU prevalence of 0.51%, a 0.10 percentage point increase over the unadjusted estimate. Larger differences were observed between adjusted and unadjusted estimates among males and younger persons.

Narcotic Overdose Mortality Adjustment Method

For the narcotic overdose correction factor, we used 2010–2016 growth in the narcotic overdose death rate to scale up self-reported IDU prevalence estimated from 2010–2011 NSDUH (Table 4). IDU prevalence in 2010–2011 was 0.24% overall and, like the more recent estimate, was higher among males, non-Hispanic whites, and young people. Overall, the ratio of narcotic overdose death in 2015–2016 compared to 2010–2011 was 1.44 and was higher among males (1.51), 18–29 year olds (1.50), and non-Hispanic blacks (1.70).

Application of the narcotic overdose death correction factor resulted in an overall adjusted IDU prevalence of 0.34%, ranging from 0.19% among Hispanic persons and non-Hispanic blacks to 0.44% among non-Hispanic whites and 0.21% among females to 0.49% among males. The overall estimate was marginally higher than the NSDUH prevalence 2010–2011 (0.24%) but lower than the unadjusted 2015–2016 estimate (0.41%).

DISCUSSION

The most recent prevalence estimates of past year IDU behavior from US household surveys range from 0.24% to 0.59% but are likely subject to multiple biases. Selection bias is likely introduced to estimates by noninclusion of unstably housed and incarcerated populations in sampling frames [47–50]. Survey nonresponse may also be differential by PWID status due to competing demands for time and fear of stigma and legal

Table 3. National Estimates of Injection Drug Use in Past 12 Months Derived From NSDUH and Adjusted Using Ratio Estimator of Injection Drug Use to 2016 Arrest Data

	Col A	Col B	Col C	Col D	Col E	Col F	Col G	Col H	Col I
Characteristic	Survey- Based Estimate for IDU in p12m for Year 2015–2016, %	Survey- Based Estimate for IDU in p12m + Arrested in p12m/ Arrested in p12m, %	Survey-Based Estimate for IDU in p12m + Not Arrested in p12m/ Not Arrested in p12m, %	Number of People Arrested in Year 2016	Number of People Not Arrested in Year 2016	(Col B × Col D)	(Col C × Col E)	Adjusted Estimate for IDU in p12m in Year 2016 (Col F + Col G)	Adjusted % IDU in p12m in Year 2016
Race									
Non-Hispanic white	0.52	8.33	0.35	NA	NA	NA	NA	NA	
Non-Hispanic black	0.17	1.23	0.12	NA	NA	NA	NA	NA	
Hispanic	0.31	5.20	0.15	NA	NA	NA	NA	NA	
Other	0.14	2.01	0.10	NA	NA	NA	NA	NA	
Sex									
Male	0.59	5.82	0.39	6 597 869	93 030 722	384 306	363 657	747 963	0.75
Female	0.23	4.91	0.15	2 471 842	98 247 406	121 390	148 747	270 136	0.27
Age group, y									
18–29	0.49	4.88	0.29	4 327 018	49 324 610	211 280	140 772	352 052	0.66
30–49	0.50	7.12	0.33	3 797 443	79 687 100	270 355	266 792	537 148	0.64
50-64	0.23		0.16	990 365	62 221 303		102 167		
Total	0.41	5.55	0.27	9 082 056	191 265 783	503 972	510 871	1 014 843	0.51

Arrest data abstracted from the 2016 Federal Bureau of Investigation Uniform Crime Report.

Abbreviations: Col, column; IDU, injection drug use; NA, not available; NSDUH, National Survey on Drug Use and Health; p12m, past 12 months.

repercussions [27, 28]. Information bias may be introduced by suboptimally designed survey items [51] and underreporting of IDU behavior [36]. Apart from the NSDUH estimate, prevalence estimates from population-based surveys were based on just 38–58 respondents, small samples that may be disproportionately affected by random error and are too small for stratification by respondent characteristics.

We observed gaps in national surveys' questionnaires that suggest their ability to measure IDU prevalence could be improved. First, survey questions should capture information about injection of prescription drugs, which has increased in recent years [3, 4], either using a universal question about any nonmedical injection or specific questions about prescription drug injection. Apart from NESARC-III, the surveys we assessed specified illicit drugs in all questionnaire items regarding injection, and NSDUH, NHANES, and NSFG items stipulated drugs that were "not prescribed." Second, questionnaire items could be harmonized across surveys to facilitate meta-analyses that combine data from small samples, as was previously carried out by Lansky and colleagues [25]. Third, specialized training should be provided to interviewers to encourage interactions with PWID that avoid stigmatizing language and attitudes and are sensitive to competing demands for time and attention [27]. Due to intensive training needs, surveys focused on substance use, such as NSDUH or NESARC, are likely better suited to measuring IDU behavior than general health surveys including limited IDU questionnaire items.

To the extent biases we identified affect estimates, they may be reduced through the use of adjustments and external data. None of the external adjustments we explored were viable using existing data but may be in the future with additional external data points. Misclassified self-reported IDU behavior may be adjusted by external estimates of self-report sensitivity, although we found very few studies outside drug treatment settings that assessed sensitivity of self-reported illicit drug use. Differences in biomarkers and assays used, including types of drugs that are detectable and over what period of time, made it impractical to compute a standardized multiplier. To our knowledge, biomarker data on drugs that are primarily injected have not been previously used to adjust survey data for PWID population size estimates. More research is needed on sensitivity of self-reported IDU, and harmonization across studies may allow findings to be combined for adjustment of survey-derived prevalence estimates.

In terms of the other 2 adjustments we explored, the arrest data adjustment method assumes persons are equally likely to underreport IDU behavior and being arrested, which is unlikely. Additionally, arrest data were not available at the person level so the number of persons arrested had to be estimated from episode-level data. Adjusting IDU prevalence using narcotic overdose mortality data may have more potential for use in the future. In our demonstration, using 2010-2016 growth in narcotic overdose mortality as an adjustment to the 2010-2011 NSDUH IDU prevalence estimate resulted in an estimate that was implausibly lower than the 2015-2016 survey-derived prevalence estimate. This adjustment in its current form is problematic because case fatality varies by drugs used or injected. Synthetic opioids, which account for a large proportion of narcotic overdose deaths in the United States, likely have higher case fatality at the same rate of injection [52, 53]. Coverage of harm reduction and overdose prevention programs also varies across time and place, which modifies the association between

Table 4. National Estimates of Injection Drug Use in Past 12 Months Derived from NSDUH and Adjusted for Temporal Change in Narcotic Overdose Death During 2010–2016

	Col A	Col B	Col C	Col D	Col E
Characteristic	Survey-Based Estimate for Year 2010–2011, %	Narcotic Overdose Death Rate per 100 000 During 2010–2011ª	Narcotic Overdose Death Rate 100 000 During 2015–2016ª	(Col C/ Col B)	Adjusted Estimate for Year 2016 (Col A × Col D), %
Race					
Non-Hispanic white	0.31	21.21	30.78	1.45	0.44
Non-Hispanic black	0.11	12.99	22.02	1.70	0.19
Hispanic	0.12	7.59	11.26	1.48	0.18
Other	0.08	3.60	4.82	1.34	0.11
Sex					
Male	0.32	21.28	32.23	1.51	0.49
Female	0.16	12.16	15.96	1.31	0.21
Age group, y					
18–29	0.36	12.81	19.24	1.50	0.55
30–49	0.31	19.69	28.39	1.44	0.44
50–64		15.88	22.46	1.41	
Total	0.24	16.69	24.06	1.44	0.34

Abbreviations: Col, column; NSDUH, National Survey on Drug Use and Health.

^aOverdose rates are calculated as (2010 death + 2011 death) / (2010 population + 2011 population) × 100 000

injection behavior and overdose mortality [54, 55], and local resource and policies (eg, medical examiner vs coroner) for death certifications may also lead to underreporting of deaths due to narcotic overdose [56]. Additionally, selection and information bias in the 2010–2011 estimate carry over to the 2015–2016 estimate, although incorporating growth in an external indicator is still likely to produce a less underestimated IDU prevalence than using the survey data alone. Data on change in types of drugs used and overdose prevention programs may increase usefulness of narcotic overdose mortality for adjusting surveyderived IDU prevalence, but due to wide geographical variation in these data points, this adjustment may be more appropriate for local versus national estimates.

Ultimately, methods apart from national surveys are needed to better estimate PWID population size. Several types of emerging methods have previously been used to estimate hardto-reach populations. Capture-recapture methods use joint probabilities of an individual appearing in multiple "captures" of a target population to estimate population size and have been carried out using both physical evidence of "captures" or appearances in multiple, linked datasets [57-65]. Service usage multipliers combine the percentage of target population members reporting usage of a particular service with the number of people receiving that service to inflate survey data to population size [51, 59, 66, 67]. Respondent-driven sampling methods have been used with successive sampling and weighting schemes to estimate the underlying population size from the size and characteristics of sampled networks [68-70]. Quantitative methods facilitating use of these techniques have advanced in recent years [71-74], but most still require intensive fieldwork and primary data collection.

Ideally, PWID population size estimates should be geographically specific enough to inform local programs but standardized across geographies to allow for comparative monitoring and optimal resource allocation. To our knowledge, standardized local estimates of PWID population size have only been estimated twice in the United States. Brady and colleagues estimated PWID population size in 96 metropolitan statistical areas through the use of an index created from external data sources anchored to previous estimates of PWID population size from 1992 and 1998 [75]. Estimates were later extended to 2007 using similar methods [76]. Current use of these estimates is limited, however, by their urban and suburban geographic focus and outdated parameters.

For future work in this area, robust analysis of large administrative data sources may offer promise for improved PWID population size estimation with fewer resource requirements compared to survey-based methods. For example, electronic medical records have been used to detect change in the frequency of conditions, such infectious endocarditis, that cause morbidity among PWID [21, 23, 24]. Additionally, injection behavior associated with narcotic overdose may be increasingly detectable from text analysis of emergency medical services, emergency room, and police response data [77]. Combining data sources may be most informative, for example using capture-recapture approaches with linked administrative data for estimation and temporal monitoring of PWID population size [60, 62, 63].

PWID are a hard-to-reach and likely growing population with complex health needs. Meaningful, population-level declines in bloodborne infections such as HIV and HCV will not be possible without increasing attention to disease surveillance and prevention in this risk group. To better serve PWID with infectious disease prevention and other health services, a much better understanding is needed of the number and characteristics of PWID in the United States, including in terms of variation across time and place. IDU prevalences derived from population surveys are subject to multiple sources of bias, and the extent of underestimation imposed by these biases is unclear. Improvements to surveys and the use of external data adjustments have potential to reduce bias in estimates, but innovative and robust nonsurveybased strategies for estimating PWID population size are urgently needed.

Notes

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