



Hepatitis C Incidence—a Comparison Between Injection and Noninjection Drug Users in New York City

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ABSTRACT *Hepatitis C virus (HCV) burdens injection drug users (IDUs) with prevalence estimated from 60–100% compared to around 5% among noninjection drug users (non-IDUs). We present preliminary data comparing the risk for HCV among IDUs and non-IDUs to inform new avenues of HCV prevention and intervention planning. Two cohorts, new IDUs (injecting ≤ 3 years) and non-IDUs (smoke/sniff heroine, crack or cocaine ≤ 10 years), ages 15–40, were street-recruited in New York City. Participants underwent risk surveys and HCV serology at baseline and 6-month follow-up visits. Person-time analysis was used to estimate annual HCV incidence. Of 683 non-IDUs, 653 were HCV seronegative, 422 returned for at least 1 follow-up visit, and 1 became HCV seropositive. Non-IDUs contributed 246.3 person-years (PY) yielding an annual incident rate of 0.4/100 PY (95% Confidence Interval [CI]=0.0–1.2). Of 260 IDUs, 114 were HCV seronegative, 62 returned for at least 1 follow-up visit, and 13 became HCV seropositive. IDUs contributed 36.3 PY yielding an annual incidence rate of 35.9/100 PY (95% CI=19.1–61.2). Among IDUs, HCV seroconverters tended to be younger (median age 25 vs. 28, respectively), and inject more frequently (61.5% vs. 34.7%, respectively) than nonseroconverters. These interim data suggest that IDUs may have engaged in high-risk practices prior to being identified for prevention services. Preventing or at least delaying transition into injection could increase opportunity to intervene. Identifying risk factors for transition into injection could inform early prevention to reduce onset of injection and risk of HCV.*

KEYWORDS *Injection drug use, Noninjection drug use, HCV incidence.*

Hepatitis C virus (HCV) continues to be highly prevalent among injection drug users (IDUs), with prevalence estimates ranging from 60% to 100%. HCV prevalence among noninjection (non-IDUs) remains low, typically around 5% in most studies.^{1–8} Reported incidence for HCV in IDUs typically ranges from 10–37/100 person-years in the United States and abroad.^{9–18} Published reports have shown high HCV incidence among recent-onset or “new” IDUs, with risk for HCV (and human immunodeficiency virus [HIV]) highest during the early stages of an injecting career,^{4,12,16,17} although some reported an increasing cumulative risk.^{19,20} This early high-risk period has been identified as occurring as early as 4 months through the first 3 years of injection drug use.^{11,12,17,21–23} This report presents preliminary data comparing the risk for

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All research was approved by the New York Academy of Medicine’s Institutional Review Board and conforms to the principles embodied in the Declaration of Helsinki.

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HCV among IDUs and non-IDUs in New York City to inform new avenues of HCV prevention and intervention planning.

In August 2000, extensive street outreach (including neighborhood mapping of high-risk neighborhoods) and screening of IDUs and non-IDUs began in Harlem and South Bronx neighborhoods of New York City, with special emphasis on enrollment of recent-onset (i.e., injecting 3 years or less before interview) or new IDUs. The majority of study participants were identified and enrolled after August 2002. Participants in the IDU cohort were eligible if they were 15–40 years of age and injected at least once in the past 2 months. Non-IDUs were eligible if they were 15–40 years of age, used heroin, crack, or cocaine 10 years or less, had no history of injection drug use or presence of stigmata, and used drugs at least once per week in the past 2 months. Study participants underwent baseline and 2-month follow-up visits over a 12-month period; these visits included risk surveys and blood draws for HCV serological testing. The institutional review boards at the New York Academy of Medicine and the New York Blood Center approved this study protocol, and informed consent was obtained from each study participant.

HCV antibodies were detected by enzyme-linked immunosorbent assay (Ortho HCV Version 2.0). Sera that were reactive on the first testing were retested in duplicate. Repeatedly reactive samples were confirmed by strip immunoblot assay (Chiron RIBA HCV 3.0 SIA). Participants returned 2 to 3 weeks later to learn their test results and receive referrals for medical care and other health and social services.

HCV seroconversion was determined by the presence of HCV antibody in previously seronegative participants. Date of HCV seroconversion was estimated to occur at the midpoint between the last seronegative visit and the first seropositive visit. Person-time analysis was used to estimate HCV incidence among cohort members who returned for follow-up.

As of August 2003, there were 683 non-IDUs tested for anti-HCV; 653 were HCV seronegative, 422 returned for at least one follow-up visit (mean follow-up time 3 months), and 1 participant became HCV seropositive. Non-IDUs contributed 246.3 person-years of follow-up time, yielding an annual incident rate of 0.4 per 100 person-years. Among 260 IDUs tested, 114 IDUs tested HCV seronegative, 62 returned for at least one follow-up visit (mean follow-up time 3 months), with 13 becoming HCV seropositive (Table 1). IDUs contributed 36.3 person-years of follow-up time, yielding an annual incidence rate of 35.9 per 100 person-years. Comparing HCV seroconverters to nonseroconverters, median age was 25 versus 28 years, respectively (Table 2). In terms of injection risk, a higher proportion of seroconverters reported high injection frequency (inject at least daily vs. less than daily) than nonseroconverters, 61.5% versus 34.7%, respectively. Among HCV-seronegative participants at baseline, at least one follow-up visit has been completed by 70% of IDUs and 73% of non-IDUs, with follow-up continuing.

TABLE 1. Comparison of HCV seroincidence rates between IDUs and non-IDUs in New York City, 2000–2003

Cohort	No. of HCV seroconverters	HCV seroconversion risk, %*	95% Confidence interval†
Non-IDU	1	0.4	(0.0–1.2)
IDU	13	35.9	(19.1–61.2)

*Calculated as number of HCV seroconverters per 100 person-years.

†The 95% confidence interval for non-IDU incidence estimate was based on a Poisson distribution; no assumption was made for the IDU incidence estimate confidence interval.

TABLE 2. Age and injection frequency of IDUs stratified by HCV seroconversion status in New York City, 2000–2003

IDU seroconversion status (N = 62)*	Median age, years (range)†	High injection frequency	
		N (%)	Odds ratio‡
HCV seroconverters	25 (23–39)	8 (61.5)	3.01
Non-HCV seroconverters	28 (17–40)	17 (34.7)	1.00

*Total number of IDUs who followed-up.

†Number (N) and proportion (%) who injected at least daily versus less than daily at baseline visit; $P < .08$.

‡95% confidence interval = 0.9–10.6.

The noteworthy finding, based on this interim analysis, is that hepatitis C incidence is dramatically higher in recent-onset IDUs than non-IDUs, even in the presence of HIV prevention efforts, which have likely contributed to declining HIV rates among IDUs.^{24–26} This suggests that, by the time an IDU has been identified for prevention services, the IDU may have already engaged in high-risk practices such as unsafe syringe use that may lead to HCV transmission. In part, this is likely because of the high efficiency of HCV transmission indicated with increased HCV risk (as compared with HIV risk) from indirect sharing practices (i.e., sharing cookers, cotton, or rinse water).^{5,27,28} Thus, the same prevention efforts that have been able to affect HIV transmission (which is less efficiently transmitted) among IDUs may not be effective in preventing the transmission of HCV.

It has also been suggested that safer injection practices, such as attending syringe-exchange programs and not sharing injection equipment, may not be employed during the start of an injecting career, suggesting that extant HIV prevention methods may be less useful in the context of HCV.^{5,29–31} Thus, the limited efficacy of extant prevention efforts coupled with the high incidence of HCV among young IDUs emphasizes the need to expand prevention efforts.

These preliminary findings must be interpreted with some caution given the small sample size and relatively short follow-up period. It is also important to note that potential sources of bias were therefore not adequately explored in this report at this stage of the study. Namely, study retention as well as external validity may have influenced the point estimates, causing over- or underestimation. Given that data presented here reflect rates approximately midway through study completion (with some fluctuation in follow-up rates), incidence rates could be unstable at this time. Although larger studies of HCV risk in new or young IDUs are needed to confirm these findings, it is not likely that the disparity in HCV risk between IDUs and non-IDUs will ease, which is the primary focus of this report.^{7,23}

These results suggest that, if injection could be prevented or at least delayed to increase opportunity to intervene with risk reduction messages, the burden of HCV among IDUs could be substantially reduced. Therefore, identifying risk factors for transition into injection could inform early prevention and intervention strategies not only to reduce injection drug use, but also to curtail risk of HCV. While prior noninjection drug use, particularly with heroin, cocaine, or crack, is a major risk factor for transition into injection drug use,^{32–34} researchers have indicated that there are subgroups of illicit drug users who do not transition into injection drug use for fear of HIV and not wanting to be identified as an injector.^{35,36} It is therefore conceivable that circumstances other than illicit noninjection drug use may be better predictors of transition into injection (e.g., high-risk social networks, lack of social

support, neighborhood characteristics, etc.). Identifying such factors so that messages can be expanded to target high-risk non-IDUs regarding the risk of injection drug use to prevent transition into injection is critical.

REFERENCES

1. Strasfeld L, Lo Y, Netski D, Thomas DL, Klein RS. The association of hepatitis C prevalence, activity, and genotype with HIV infection in a cohort of New York City drug users. *J Acquir Immune Defic Syndr*. 2003;33:356–364.
2. Habib SE, Lovejoy FH, Aspin C. Hepatitis C prevalence and risk behavior of injecting drug users in Sydney: a continuing concern. *Southeast Asian J Trop Med Public Health*. 2001;32:823–834.
3. Murrill CS, Weeks H, Castrucci BC, et al. Age-specific seroprevalence of HIV, hepatitis B virus, and hepatitis C virus infection among injection drug users admitted to drug treatment in six US cities. *Am J Public Health*. 2002;92:385–387.
4. Garfein RS, Vlahov D, Galai N, Doherty MC, Nelson KE. Viral infections in short-term injection drug users: the prevalence of the hepatitis C, hepatitis B, human immunodeficiency, and human T-lymphotropic viruses. *Am J Public Health*. 1996;86:655–661.
5. Garfein RS, Doherty MC, Monterroso ER, Thomas DL, Nelson KE, Vlahov D. Prevalence and incidence of hepatitis C virus infection among young adult injection drug users. *J Acquir Immune Defic Syndr Hum Retrovirol*. 1998;18(suppl 1):S11–S19.
6. Thomas DL, Vlahov D, Solomon L, et al. Correlates of hepatitis C virus infections among injection drug users. *Medicine (Baltimore)*. 1995;74:212–220.
7. Quaglio G, Lugoboni F, Pajusco B, et al. Factors associated with hepatitis C virus infection in injection and noninjection drug users in Italy. *Clin Infect Dis*. 2003;37:33–40.
8. Tortu S, Neaigus A, McMahon J, Hagen D. Hepatitis C among noninjecting drug users: a report. *Subst Use Misuse*. 2001;36:523–534.
9. Van den Hoek JA, Van Haastrecht HJ, Goudsmit J, de Wolf F, Coutinho RA. Prevalence, incidence, and risk factors of hepatitis C virus infection among drug users in Amsterdam. *J Infect Dis*. 1990;162:823–826.
10. Chamot E, de Saussure P, Hirschel B, Deglon JJ, Perrin LH. Incidence of hepatitis C, hepatitis B and HIV infections among drug users in a methadone-maintenance programme. *AIDS*. 1992;6:430–431.
11. van Ameijden EJ, Van den Hoek JA, Mientjes GH, Coutinho RA. A longitudinal study on the incidence and transmission patterns of HIV, HBV and HCV infection among drug users in Amsterdam. *Eur J Epidemiol*. 1993;9:255–262.
12. Garfein RS, Doherty MC, Monterroso ER, Thomas DL, Nelson KE, Vlahov D. Prevalence and incidence of hepatitis C virus infection among young adult injection drug users. *J Acquir Immune Defic Syndr Hum Retrovirol*. 1998;18(suppl 1):S11–S19.
13. van Beek I, Dwyer R, Dore GJ, Luo K, Kaldor JM. Infection with HIV and hepatitis C virus among injecting drug users in a prevention setting: retrospective cohort study. *BMJ*. 1998;317:433–437.
14. Villano SA, Vlahov D, Nelson KE, Lyles CM, Cohn S, Thomas DL. Incidence and risk factors for hepatitis C among injection drug users in Baltimore, Maryland. *J Clin Microbiol*. 1997;35:3274–3277.
15. Rezza G, Saggiocca L, Zaccarelli M, Nespoli M, Siconolfi M, Baldassarre C. Incidence rate and risk factors for HCV seroconversion among injecting drug users in an area with low HIV seroprevalence. *Scand J Infect Dis*. 1996;28:27–29.
16. Miller CL, Johnston C, Spittal PM, et al. Opportunities for prevention: hepatitis C prevalence and incidence in a cohort of young injection drug users. *Hepatology*. 2002;36:737–742.
17. Hagan H, Thiede H, Weiss NS, Hopkins SG, Duchin JS, Alexander ER. Sharing drug preparation equipment as a risk factor for hepatitis C. *Am J Public Health*. 2001;91:42–46.
18. Hahn JA, Page-Shafer K, Lum PJ, et al. Hepatitis C virus seroconversion among young injection drug users: relationships and risks. *J Infect Dis*. 2002;186:1558–1564.

19. Diaz T, Des Jarlais DC, Vlahov D, et al. Factors associated with prevalence hepatitis C: differences among young adult injection drug users in lower and upper Manhattan, New York City. *Am J Public Health*. 2001;91:23–30.
20. Thorpe LE, Ouellet LJ, Levy JR, Williams IT, Monterroso ER. Hepatitis C virus infection: prevalence, risk factors, and prevention opportunities among young injection drug users in Chicago, 1997–1999. *J Infect Dis*. 2000;182:1588–1594.
21. Nelson KE, Vlahov D, Solomon L, Cohn S, Munoz A. Temporal trends of incident human immunodeficiency virus infection in a cohort of injecting drug users in Baltimore, Maryland. *Arch Intern Med*. 1995;155:1305–1311.
22. Nicolosi A, Leite ML, Musicco M, Molinari S, Lazzarin A. Parenteral and sexual transmission of human immunodeficiency virus in intravenous drug users: a study of seroconversion. The Northern Italian Seronegative Drug Addicts (NISDA) study. *Am J Epidemiol*. 1992;135:225–233.
23. Chang CJ, Lin CH, Lee CT, Chang SJ, Ko YC, Liu HW. Hepatitis C virus infection among short-term intravenous drug users in southern Taiwan. *Eur J Epidemiol*. 1999;15:597–601.
24. Monterroso ER, Hamburger ME, Vlahov D, et al. Prevention of HIV infection in street-recruited injection drug users. The Collaborative Injection Drug User Study (CIDUS). *J Acquir Immune Defic Syndr*. 2000;25:63–70.
25. Centers for Disease Control and Prevention and HIV/AIDS Prevention Research Synthesis Project. *Compendium of HIV Prevention Interventions with Evidence of Effectiveness*. August 2001:1–64.
26. Semaan S, Des J, Sogolow E, et al. A meta-analysis of the effect of HIV prevention interventions on the sex behaviors of drug users in the United States. *J Acquir Immune Defic Syndr*. 2002;30(suppl 1):S73–93.
27. Hagan H, Des J. HIV and HCV infection among injecting drug users. *Mt Sinai J Med*. 2000;67:423–428.
28. Grund JP, Friedman SR, Stern LS, et al. Syringe-mediated drug sharing among injecting drug users: patterns, social context and implications for transmission of blood-borne pathogens. *Soc Sci Med*. 1996;42:691–703.
29. Doherty MC, Garfein RS, Monterroso E, Brown D, Vlahov D. Correlates of HIV infection among young adult short-term injection drug users. *AIDS*. 2000;14:717–726.
30. Carneiro M, Fuller C, Doherty MC, Vlahov D. HIV prevalence and risk behaviors among new initiates into injection drug use over the age of 40 years old. *Drug Alcohol Depend*. 1999;54:83–86.
31. Fuller CM, Vlahov D, Latkin CA, Ompad DC, Celentano DD, Strathdee SA. Social circumstances of initiation of injection drug use and early shooting gallery attendance: implications for HIV intervention among adolescent and young adult injection drug users. *J Acquir Immune Defic Syndr*. 2003;32:86–93.
32. van Ameijden EJ, Van den Hoek JA, Hartgers C, Coutinho RA. Risk factors for the transition from noninjection to injection drug use and accompanying AIDS risk behavior in a cohort of drug users. *Am J Epidemiol*. 1994;139:1153–1163.
33. Irwin KL, Edlin BR, Faruque S, et al. Crack cocaine smokers who turn to drug injection: characteristics, factors associated with injection, and implications for HIV transmission. The Multicenter Crack Cocaine and HIV Infection Study Team. *Drug Alcohol Depend*. 1996;42:85–92.
34. Fuller CM, Vlahov D, Arria AM, Ompad DC, Garfein R, Strathdee SA. Factors associated with adolescent initiation of injection drug use. *Public Health Rep*. 2001;116(suppl 1):136–145.
35. Des J, Perlis T, Friedman SR, et al. Declining seroprevalence in a very large HIV epidemic: injecting drug users in New York City, 1991 to 1996. *Am J Public Health*. 1998;88:1801–1806.
36. Ouellet LJ, Wiebel WW, Jimenez AD. Team research methods for studying intranasal heroin use and its HIV risks. *NIDA Res Monogr*. 1995;157:182–211.