Hepatitis Elimination in Persons who Inject Drugs

Danae (Dee) Bixler, MD, MPH Division of Viral Hepatitis Centers for Disease Control and Prevention





Disclaimer

The findings and conclusions in this presentation are those of the author and do not necessarily represent the official position of the Centers for Disease Control and Prevention.



Injection Drug Use

2011 US Estimate:

- Lifetime prevalence
 - 2.6% persons > 13 years
 - 6.6 million persons
- Past year prevalence
 - 0.30%
 - 774,434
- 2011 prevalence
 - 2.2% HIV (+)
 - 43.1% anti-HCV (+)

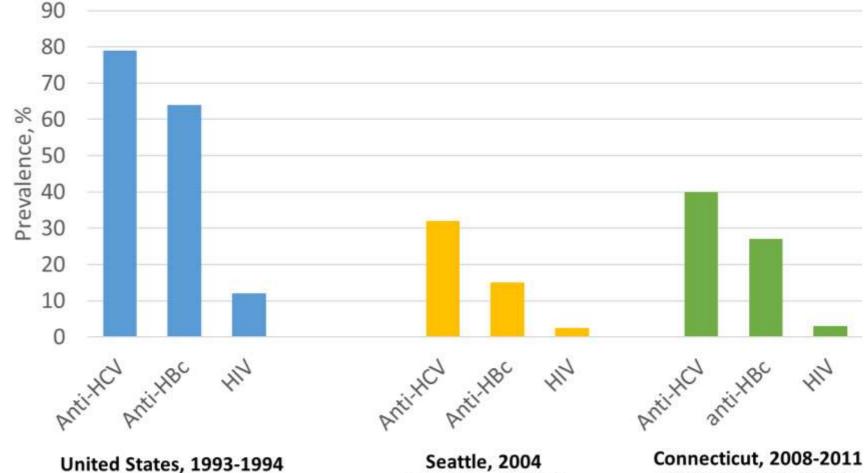


Lansky (2014) PLoS One 9(5): e97596. Degenhardt (2017)The Lancet. Global Health 5(12): e1192-e1207.

2008-2017 Global Estimate:

- 10·2–23·7 million persons 15– 64 years
- 17·8% HIV (+)
- 52·3% anti-HCV (+)

HIV, Anti-HBc and Anti-HCV Seroprevalence in Persons Who Inject Drugs, Selected US Studies



Murill CS, et al (2002) AJPH, 92: 385.

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Burt RD, et al (2007) J Urban Health, 84(3): 436 Connecticut, 2008-2011 Akselrod H, et al (2014) AJPH, 104:1713

Anti-HCV and HCV RNA Prevalence in Substance Use Treatment

Reference	Year	Location	Population	Anti- HCV	HCV- RNA
Norton, J Subst Abuse Treat 75: 38	2017	Bronx, NY	Primary care patients initiating buprenorphine	52.9%	33.9%
Carey, J Subst Abuse Treat 66: 54	2016	Boston, MA	Primary care patients receiving buprenorphine	47.7%	32.3%
Jordan. Drug Alcohol Depend 152: 194	2015	New York City	Methadone maintenance treatment programs (MMTP)	67%	
Perlman, J Addict Dis 33: 322	2014	New York, San Francisco	MMTP	59%	
Martinez, J Viral Hepat 19: 47	2012	New York City San Diego	MMTP	64.1%	39.2%



Hepatitis B and C



Symptoms of acute hepatitis

Systemic

- Fever
- Fatigue
- Loss of appetite
- Joint pain

Gastrointestinal

- Nausea
- Vomiting
- Abdominal pain
- Diarrhea

Jaundice

- Yellow skin
 and eyes
- Dark urine
 - Clay-colored bowel movements



Asymptomatic

Hepatitis B
■Most children < 5 years</p>
■Immunosuppressed adults
■50-70% of persons ≥ 5 years

<u>Hepatitis C</u> □ 70-80%



Chronic infection Hepatitis B 90% of infants 25%-50% of children 1-5 years of age 5% of adults

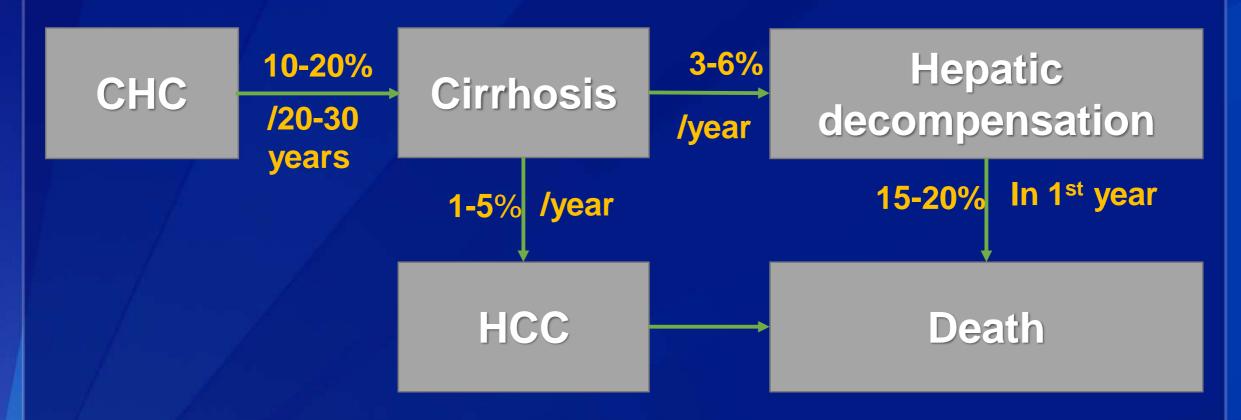
Hepatitis C 75%–85% of cases



Natural history of chronic hepatitis B (CHB) Inflammation, elevated ALT □Liver failure □40-50% of deaths are liver-related at average 60 years



Natural history of chronic hepatitis C (CHC)





Westbrook, R.H. and G. Dusheiko, Natural history of hepatitis C. J Hepatol, 2014. 61(1 Suppl): p. S58-68.

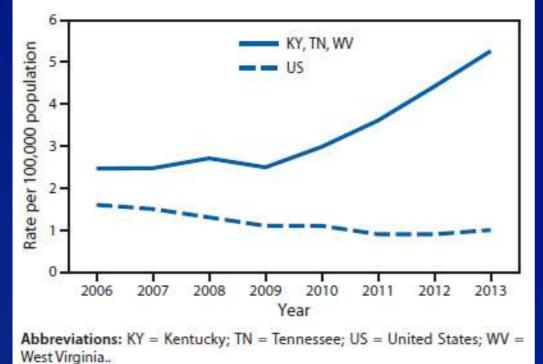
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Surveillance Data

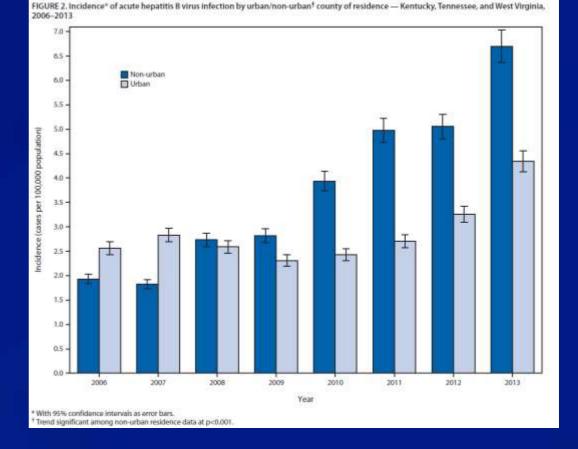


Hepatitis B in PWID

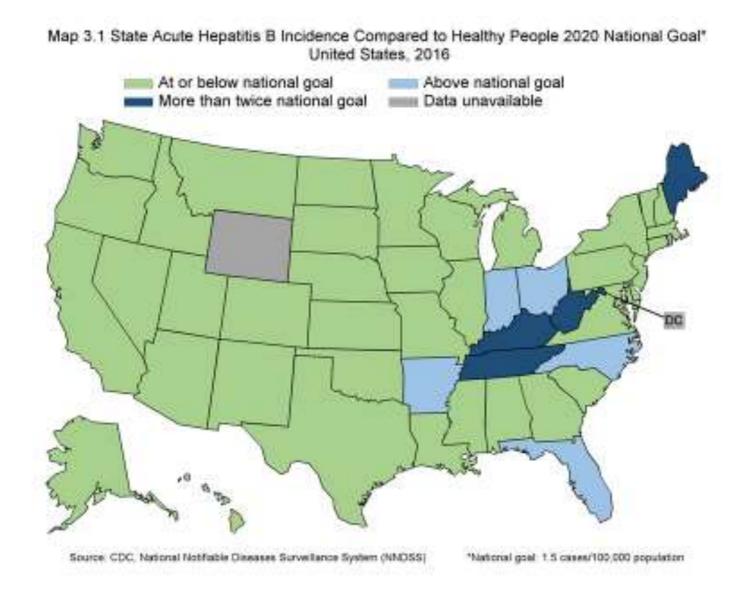
FIGURE 1. Incidence of acute hepatitis B virus infection, by year— United States and Kentucky, Tennessee, and West Virginia, 2006–2013



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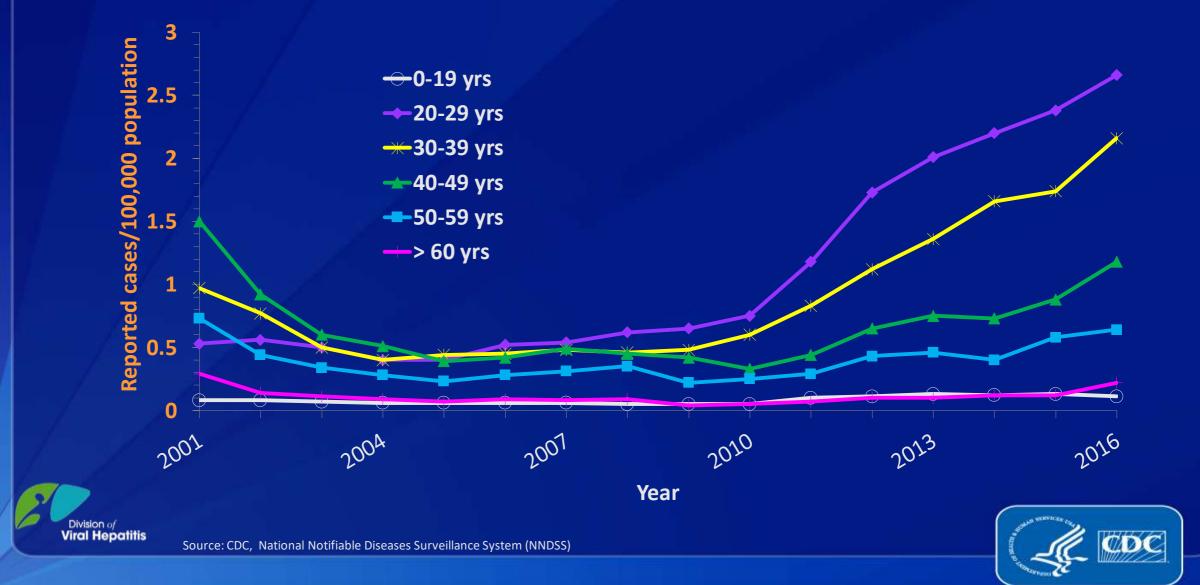


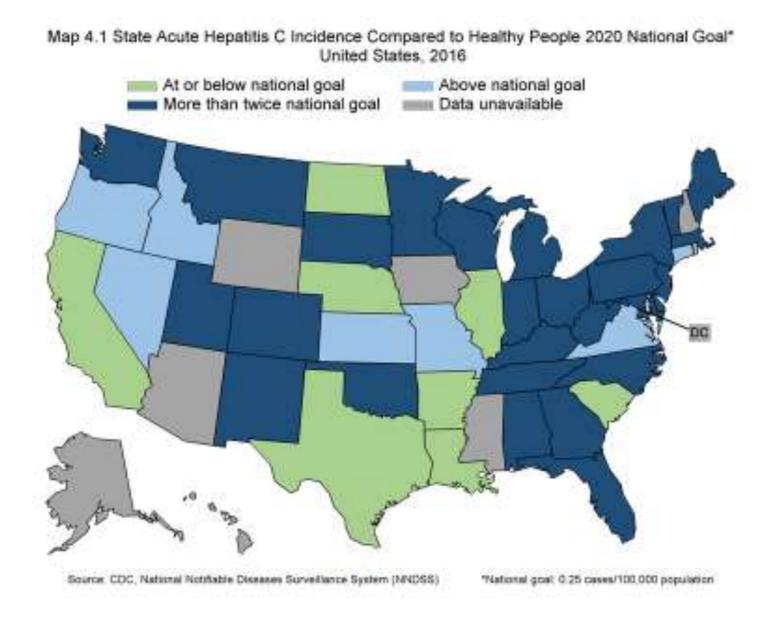
https://www.cdc.gov/mmwr/volumes/65/wr/pdfs/mm6503a2.pdf



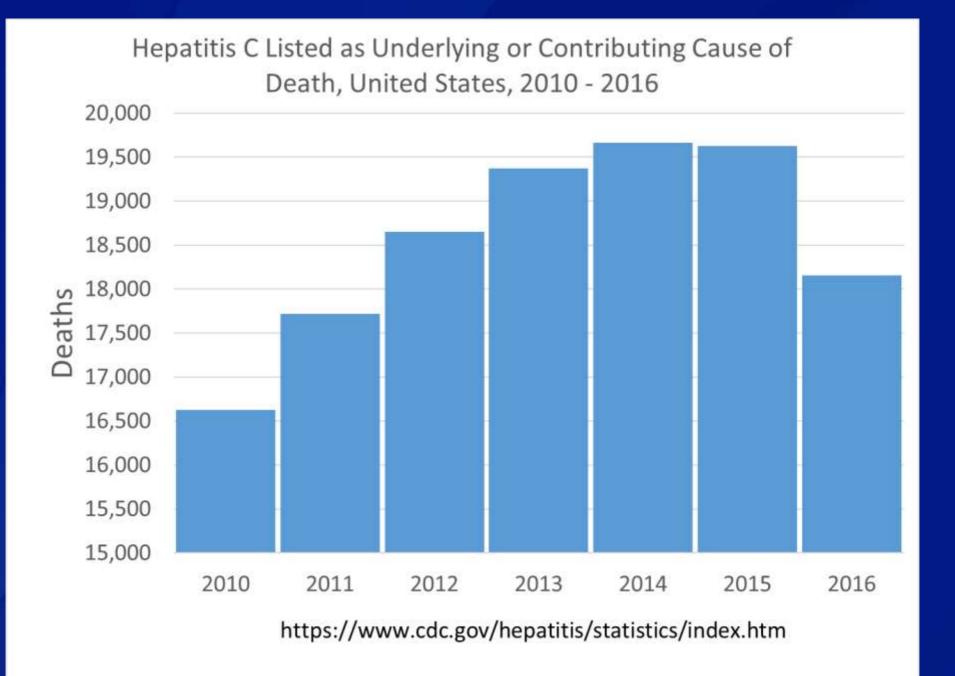


Incidence of acute hepatitis C, by age group — United States, 2001–2016







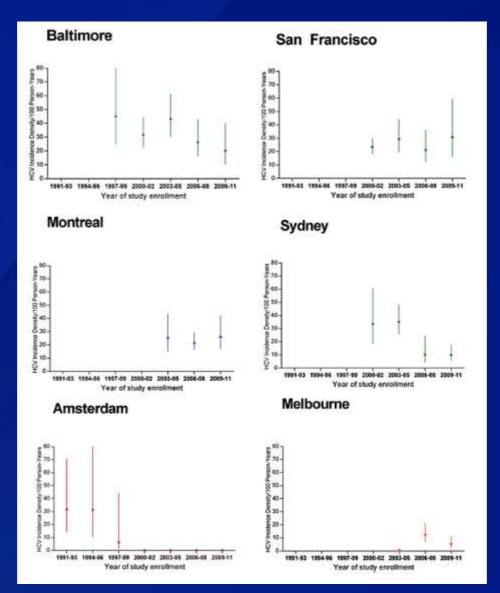




Hepatitis B and C Elimination in PWID



Trends in Hepatitis C Virus (HCV) per 100 Person-Years for PWID Across Calendar Period, by City



Morris, CID, 2017; 64:860



Combined MAT (OST) and SSPs (NSP) Decreases HCV Transmission by 74%

Figure 5. Forest plot of comparison: 4 Combined OST and high/low NSP versus no OST and low/no NSP, outcome: 4.1 HCV incidence adjusted analyses.

			anti-HCV negative	new HCV cases		Risk Ratio	Risk Ratio
Study or Subgroup	log[Risk Ratio]	SE	Tota	Total	Weight	IV, Random, 95% CI	IV, Random, 95% Cl
12.1.1 High NSP coverage							12.221
Bruneau 2015 (pers comm)	-0.52763	0.27035	183	102	24.7%	0.59 [0.35, 1.00]	
Palmateer 2014a	-2.99573	0.73734	2396	392	13.5%	0.05 [0.01, 0.21]	
Van Den Berg 2007	-1.02165	0.528005	151	17	18.2%	0.36 [0.13, 1.01]	
Subtotal (95% CI)			2730	511	56.4%	0.26 [0.07, 0.89]	
Heterogeneity: Tau# = 0.94; C	hi# = 9.99, df = 2 (P = 0.007); I	°= 80%				
Test for overall effect: Z = 2.14	(P = 0.03)						
12.1.2 Low NSP coverage							
Palmateer 2014a	-0.52763	0.420199	2396	392	20.9%	0.59 [0.26, 1.34]	
Van Den Berg 2007	0.157004	0.348184	123	45	22.7%	1.17 [0.59, 2.32]	
Subtotal (95% CI)			2519	437	43.6%	0.87 [0.44, 1.68]	-
Heterogeneity: Tau ^a = 0.09; C	hi ² = 1.57, df = 1 (P = 0.21); I ²	= 36%				
Test for overall effect: Z = 0.43	B (P = 0.67)						
Total (95% CI)			5249	948	100.0%	0.45 [0.22, 0.94]	-
Heterogeneity, Tau ² = 0.50; C	hi ² = 15.93, df = 4	(P = 0.003)	I ² = 75%				ta di di di
Test for overall effect: Z = 2.12	아님은 전 것은 아이에 가지 않는 것 같아요. 아이가 있다.		100 mm - 1997 - 1997 - 1997 - 1997 - 1997 - 1997 - 1997 - 1997 - 1997 - 1997 - 1997 - 1997 - 1997 - 1997 - 199 - 1997 - 1997 - 1997 - 1997 - 1997 - 1997 - 1997 - 1997 - 1997 - 1997 - 1997 - 1997 - 1997 - 1997 - 1997 - 1997				0.01 0.1 1 10 100
Test for subgroup differences	1 C	1 (P = 0.09)	. I² = 64.9%				Favours combined OST-NSP Favours no OST-NSP



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What are comprehensive syringe services programs (SSPs)?

- Sterile needles, syringes, injection supplies
- Safe disposal of injection supplies
- Vaccinations, PrEP
- Testing for hepatitis, HIV, referral to treatment
- Naloxone and training
- STD prevention and treatment
- Patient-centered reproductive health including LARC
- Linkage to MAT



Treatment of Hepatitis C in PWID
Comparable SVR in PWID
~ 100% for persons retained in follow-up
LTFU as high as 20% for persons in unstable housing
Reinfection rate ≈2 per 100 person-years
Increased in IDU relapse

Aspinall EJ, CID, 2013; 57 (Suppl 2): S80-S89. Lalezari J, J Hepatol, 2015; 63:364-369. Ho SB, Clin Gastroenterol Hepatol, 2015; 13: 2004-2014. Dore GJ, Ann Intern Med, 2016; 165: 625-634. Morris L. Int J Drug Pol, 2017; 47: 216-220. Read P. Int J Drug Pol, 2017; 47: 209-215.
Simmons B, CID, 2016; 62: 683-94. Midgard H. J Hepatol, 2016; 54: 1020-1026. Martinello N. J Viral Hepat, 2017; 24: 359=370. Weir A. Drug Alcohol Dep, 2016; 165:



HCV treatment integrated into substance use treatment

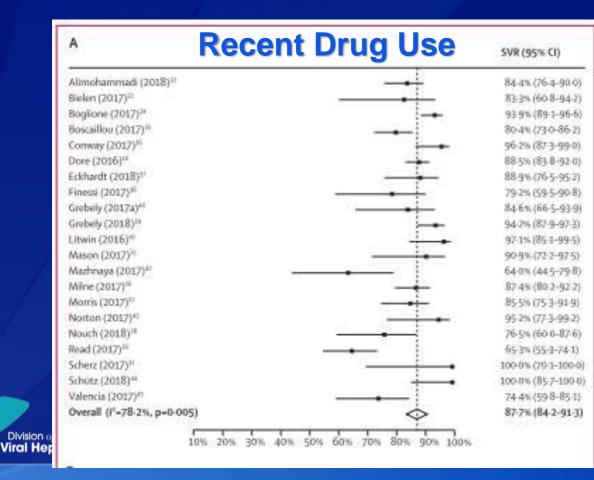
First consecutive 75 CHC patients opting to initiate DAAs 2013-2015.

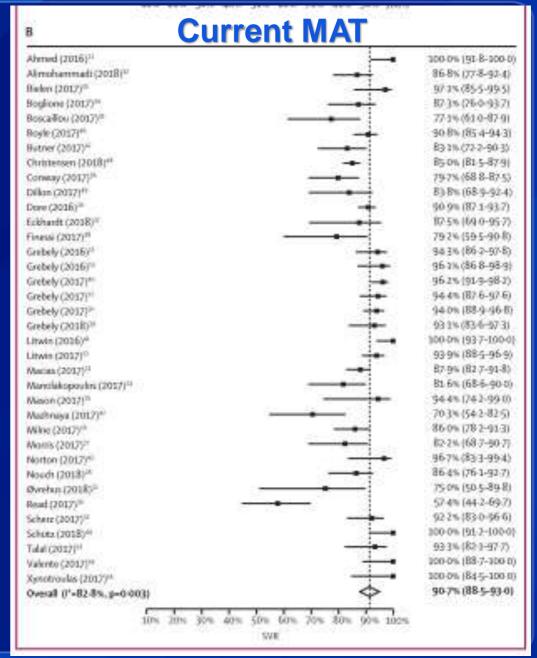
10 LTFU

- 98% SVR among those completing therapy
- 23% ongoing illicit drug use

Meta-Analysis – SVR Among PWID Treated with DAAs for HCV

Hajarizadeh B, et al. (2018). Lancet Gastroenterol Hepatol 3: 754.





Randomized Trial for HCV Treatment Strategies in Methadone Maintenance Treatment Clinics

		Self- administered Treatment	
HCV RNA (+) persons on methadone	Randomized Stratified by: IL28B genotype	Modified directly- observed treatment	
	HIV status Cirrhosis vs. no cirrhosis	Group treatment	

Outcomes

- Adherence
- SVR
- Toxicology
- Psychosocial metrics
- Costs

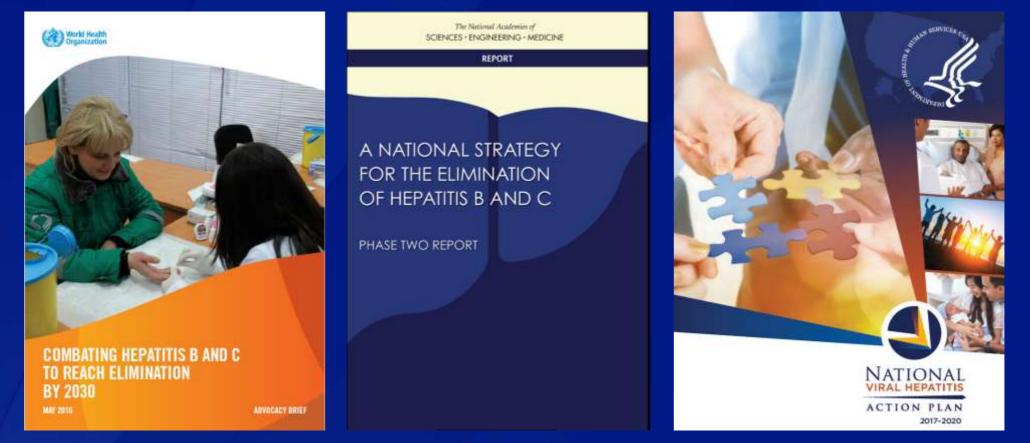


Akiyama MJ, BMC Infect Dis 2018; 18: 74. ClinicalTrials.gov (NCT01857245), May 20th, 2013.

Elimination Strategies



Elimination of Hepatitis B and C



www.who.int/hepatitis/publications/hep-elimination-by-2030-brief/en/

http://www.nationalacademies.org/hmd/reports/2017/national-strategy-for-the-elimination-of-hepatitis-b-and-c.aspx www.hhs.gov/sites/default/files/National%20Viral%20Hepatitis%20Action%20Plan%202017-2020.pdf

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Selected HCV Indicators, National Viral Hepatitis Action Plan, 2017 - 2020

Indicator	Baseline: 2014	2020 Goal
Decrease new HCV infections by 60%; estimated (reported)	30,500 (2,194)	10,889 (783)
Reduce HCV-related deaths by 25%	18,659	14,744
Decrease new HCV infections among persons 20-39 years by 60%	1,561	624



National Viral Hepatitis Action Plan, 2017 – 2020, Selected Strategies Addressing Injection Drug Use

- 1. Screen for substance use disorder and offer:
 - HAV and HBV vaccination
 - HBV and HCV testing
 - Assessment for viral hepatitis treatment if infected
 - Referral to MAT, SSPs
- 2. Ensure PWID have access to evidence-based treatment services
- 3. Expand access/delivery of hepatitis prevention and treatment services in correctional settings

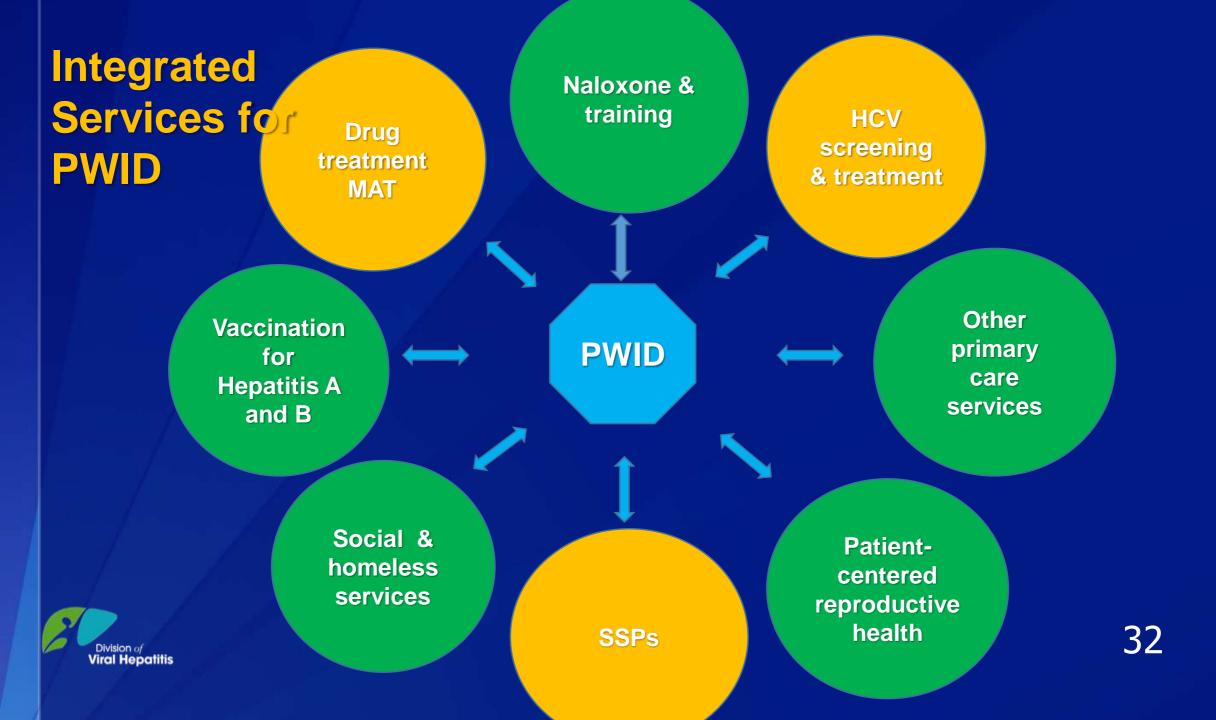


Recommendations for Screening and Treatment of HCV Infection in People Who Inject Drugs (PWID)					
RECOMMENDED	RATING 🖸				
Annual HCV testing is recommended for PWID with no prior testing, or past negative testing and subsequent injection drug use. Depending on the level of risk, more frequent testing may be indicated.	lla, C				
Substance use disorder treatment programs and needle/syringe exchange programs should offer routine, opt-out HCV-antibody testing with reflexive or immediate confirmatory HCV-RNA testing and linkage to care for those who are infected.	lla, C				
PWID should be counseled about measures to reduce the risk of HCV transmission to others.	I, C				
PWID should be offered linkage to harm reduction services when available, including needle/syringe service programs and substance use disorder treatment programs.	I, B				
Active or recent drug use or a concern for reinfection is not a contraindication to HCV treatment.	lla, B				

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IDSA / AASLD Recommendations for PWID Annual testing for HCV Routine, opt-out HCV testing and linkage to care: SUD treatment facilities SSPs Counsel about measures to reduce risk □ Offer linkage to: SSPs SUD treatment Active or recent drug use or concern for reinfection is not a contraindication to HCV treatment.





Thankyou Danae (Dee) Bixler nqd0@cdc.gov (404) 718-3208



Extra Slides



"Call to action"

<u>Action Step 1:</u> Implement screening for OUD in all relevant health care settings.

<u>Action Step 2:</u> For patients with positive screening results, immediately prescribe effective medication for OUD and/or opioid withdrawal symptoms.

<u>Action Step 3:</u> Develop hospital-based protocols that facilitate OUD treatment initiation and linkage to community-based treatment upon discharge.

<u>Action Step 4:</u> Hospitals, medical schools, physician assistant schools, nursing schools, and residency programs should increase training to identify and treat OUD.

<u>Action Step 5:</u> Increase access to addiction care and funding to states to provide effective medications to treat OUD.



Myth	Reality	Possible Policy Response
Buprenorphine treatment is more dangerous than oth- er chronic disease man- agement.	Buprenorphine treatment is simpler than many other routine treatments in primary care, such as titrating insulin or starting anticoagulation. But physicians receive little training in it.	Amend federal buprenorphine-treatment eligibili- ty requirements to include training completed during medical school and require training during medical school or residency. Add com- petency questions to U.S. Medical Licensing Examination and other licensing exams.
Use of buprenorphine is sim- ply a "replacement" addic- tion.	Addiction is defined as compulsively using a drug despite harm. Taking a prescribed medication to manage a chronic illness does not meet that definition.	Public health campaign to reduce stigma associ- ated with addiction treatment, similar to past campaigns (e.g., HIV) that provided educa- tion and challenged common myths.
Detoxification for opioid use disorder is effective.	There are no data showing that detoxifica- tion programs are effective at treating opioid use disorder. In fact, these inter- ventions may increase the likelihood of overdose death by eliminating tolerance.	Advocacy from professional physician organiza- tions to educate federal and state agencies and policymakers about evidence-based treat- ment and the lack of evidence for short-term "detoxification" treatment.
Prescribing buprenorphine is time consuming and bur- densome.	Treating patients with buprenorphine can be uniquely rewarding. In-office inductions and intensive behavioral therapy are not required for effective treatment.	Develop and disseminate protocols for primary care settings that emphasize out-of-office in- duction and treatment.
Reducing opioid prescribing alone will reduce overdose deaths.	Despite decreasing opioid prescribing, over- dose mortality has increased. Patients with opioid use disorder may shift to the illicit drug market, where the risk of over- dose is higher.	Develop a national system of virtual consultation for physicians to reach addiction and pain specialists who can support treatment of pa- tients with suspected opioid use disorder.

Wakeman, NEJM, 2018; 379(1): 1-4.

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P.1-1	-
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I	Evidence and/or general agreement that a given diagnostic evaluation, procedure, or treatment is beneficial, useful, and effective.
Ш	Conflicting evidence and/or a divergence of opinion about the usefulness and efficacy of a diagnostic evaluation, procedure, or treatment.
lla	Weight of evidence and/or opinion is in favor of usefulness and efficacy.
llb	Usefulness and efficacy are less well established by evidence and/or opinion.
Ш	Conditions for which there is evidence and/or general agreement that a diagnostic evaluation, procedure, or treatment is not useful and effective or if it in some cases may be harmful.

	Level
А	Data derived from multiple randomized clinical trials, meta-analyses, or equivalent.
В	Data derived from a single randomized trial, nonrandomized studies, or equivalent.
с	Consensus opinion of experts, case studies, or standard of care.

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Injection Drug Use Associations

- Hepatitis A
- Hepatitis B
- Hepatitis C
- HIV
- MRSA and other bacterial infections
- Perinatal infection HBV, HCV, HIV



HCV Seroprevalence in Emergency Departments

Study	Year	Location	Population	Anti- HCV	HCV RNA
Cornett, OFID 5(4): ofy065.	2018	Rutgers, NJ	Boomers	6.6%	2.4%
Hsieh, CID 62:1059	2016	Baltimore	All patients with blood draws	13.8%	
Lyons, CID 62:1066	2016	Cincinnati	18-64	13.9%	11.1%
Franco, OFID3(4): ofw211	2016	Birmingham	Boomers	11%	7.6%
White, Ann Emerg Med 67: 119	2016	Oakland, CA	Boomers, PWID	10.3%	8.9%



Anti-HCV and HCV RNA Prevalence in Corrections

Study	Year	Location	Population	Setting	Anti- HCV	HCV RNA
De la Flor Public Health Rep 132:617	2017	Dallas	Universal, opt-out	jail	16.4%	
Akiyama, Public Health Rep 132: 41	2017	New York City	Boomers, risk	jail	20.6%	
Hawks, J Viral Hepat 23:473	2016	Bronx, NY	Not specified	Post- release	33%	20.6%
Mahowald, J Correct Health Care 22: 41	2016	Pennsylvania	> 9 years	State system	18.1%	5.2%
Schoenbachler, Public Health Rep 131 Suppl 2: 98	2016	North Carolina South Carolina	All At risk / all	jails	13.2% 8.0%	9.8% 4.0%
Stockman, Public Health Rep 131: 544	2016	Wisconsin	<u>></u> 18 years	prison	12.5%	8.9%