U.S. Preventive Services Task Force

Screening for Hepatitis C Virus Infection in Adolescents and Adults: **Recommendation Statement**

Summary of Recommendation and Evidence

The USPSTF recommends screening for hepatitis C virus (HCV) infection in adults aged 18 to 79 years (Table 1). B recommendation.

See *Table 1* for a more detailed summary of the recommendation for clinicians.

Importance

HCV is the most common chronic blood-borne pathogen in the United States and is a leading cause of complications from chronic liver disease.1 HCV infection is associated with more deaths than the top 60 other reportable infectious diseases combined, including HIV.2 The most important risk factor for HCV infection is past or current injection drug use. In the United States, an estimated 4.1 million persons have past or current HCV infection (i.e., they test positive for the anti-HCV antibody). Of these persons who test positive for the anti-HCV antibody, approximately 2.4 million have current infections based on testing with molecular assays for HCV RNA.^{1,3-5} The estimated prevalence of chronic HCV infection is approximately 1.0% (2013 to 2016).6 An estimated 44,700 new HCV

See related Putting Prevention into Practice on page 367. As published by the USPSTF.

This summary is one in a series excerpted from the Recommendation Statements released by the USPSTF. These statements address preventive health services for use in primary care clinical settings, including screening tests, counseling, and preventive medications.

The complete version of this statement, including supporting scientific evidence, evidence tables, grading system, members of the USPSTF at the time this recommendation was finalized, and references, is available on the USPSTF website at https://www.uspreventiveservicestaskforce.org/.

This series is coordinated by Kenny Lin, MD, MPH, deputy editor.

A collection of USPSTF recommendation statements published in AFP is available at https://www.aafp.org/afp/uspstf. infections occurred in the United States in 2017.7 Cases of acute HCV infection have increased approximately 3.8-fold (2010 to 2017) over the last decade because of increasing injection drug use and improved surveillance.7 The most rapid increase in acute HCV incidence has been in young adults aged 20 to 39 years who inject drugs, with increases in both sexes but more pronounced in men.⁷ Rates increased especially in American Indian/Alaska Native and non-Hispanic white populations.⁷

Assessment of Magnitude of Net Benefit

The USPSTF concludes with moderate certainty that screening for HCV infection in adults aged 18 to 79 years has substantial net benefit.

See Table 1 and Table 2 for more information. on the USPSTF recommendation rationale and assessment. For more details on the methods the USPSTF uses to determine net benefit, see the USPSTF Procedure Manual.8

Practice Considerations

PATIENT POPULATION UNDER **CONSIDERATION**

This recommendation applies to all asymptomatic adults aged 18 to 79 years without known liver disease.

ASSESSMENT OF RISK

Although all adults aged 18 to 79 years should be screened, a number of risk factors increase risk. The most important risk factor for HCV infection is past or current injection drug use. In the United States, recent increases in HCV incidence have predominantly been among young persons who inject drugs.^{1,9} Approximately one-third of persons who inject drugs aged 18 to 30 years are infected with HCV, and 70% to 90% of older persons who inject drugs are infected.9 Clinicians may want to consider screening in adolescents younger than 18 years and in adults older than

79 years who are at high risk (e.g., past or current injection drug use).

Pregnant adults should be screened. HCV prevalence has doubled in women aged 15 to 44 years from 2006 to 2014. 1,10,11 From 2011 to 2014, 0.73% of pregnant women tested had an HCV infection, with a 68% increase in the proportion of infants born to HCV-infected mothers. 1,10 Approximately 1,700 infected infants are born annually to 29,000 HCV-infected mothers. 1,11 Because of the increasing prevalence of HCV in women aged 15 to 44 years and in infants born to HCV-infected mothers, clinicians may want to consider screening pregnant persons younger than 18 years.

SCREENING TESTS

Screening with anti-HCV antibody testing followed by polymerase chain reaction testing for HCV RNA is accurate for identifying patients with chronic HCV infection. Currently, diagnostic evaluations are often performed with various noninvasive tests that have lower risk for harm than liver biopsy for diagnosing fibrosis stage or cirrhosis in persons who screen positive. 12

Among patients with abnormal results on liver function tests (measurement of aspartate aminotransferase, alanine aminotransferase, or bilirubin levels) who were tested for reasons other than HCV screening, finding the cause of the

TABLE 1 Screening for HCV Infection in Adolescents and Adults: Clinical Summary of the USPSTF Recommendation What does the USPSTF For adults aged 18 to 79 years: Grade B recommend? Screen adults for HCV infection. To whom does this rec-Asymptomatic adults aged 18 to 79 years (including pregnant persons) without known liver ommendation apply? disease. What's new? This recommendation expands the population that should be screened. The USPSTF now recommends that all adults aged 18 to 79 years be screened. Previously, it recommended screening adults born between 1945 and 1965 and others at high risk. How to implement this Screen. Screen adults aged 18 to 79 years with anti-HCV antibody testing followed by confirmarecommendation? tory polymerase chain reaction testing. a. The USPSTF also suggests that clinicians consider screening persons younger than 18 years and older than 79 years who are at high risk for infection (e.g., those with past or current injection drug use). Adults with a positive screening test result are usually followed up with a diagnostic evaluation using 1 of various noninvasive tests. Treatment typically consists of oral direct-acting antiviral regimens for 8 to 12 weeks. Important considerations include: · Communicating that screening is voluntary and undertaken only with the patient's knowledge • Informing patients about HCV infection, how it can (and cannot) be acquired, the meaning of positive and negative test results, and the benefits and harms of treatment • Providing patients the opportunity to ask questions and to decline screening How often? One-time screening for most adults. Periodically screen persons with continued risk for HCV infection (e.g., persons with past or current injection drug use). There is limited evidence to determine how often to screen persons at increased risk. What are other The USPSTF has made recommendations on screening for hepatitis B virus infection in pregnant relevant USPSTF persons, hepatitis B virus infection in adults, and HIV infection. These recommendations are availrecommendations? able at www.uspreventiveservicestaskforce.org. Where to read the Visit the USPSTF website to read the full recommendation statement. This includes more details full recommendation on the rationale of the recommendation, including benefits and harms; supporting evidence; and statement? recommendations of others. Note: For a summary of the evidence systematically reviewed in making this recommendation, the full recommendation statement, and supporting documents, go to https://www.uspreventiveservicestaskforce.org/. HCV = hepatitis C virus; USPSTF = U.S. Preventive Services Task Force.

Summary of USPSTF Rationale for Screening for HCV Infection	
Population	Assessment
Detection	There is adequate evidence that HCV testing (screening for the anti-HCV antibody followed by confirmation of active infection by HCV RNA assay for persons who test positive) accurately detect HCV infection.
	There is adequate evidence for 1-time testing in all adults and periodic testing in persons at continued risk of new HCV infection.
	There is inadequate evidence on the timing of repeat testing.
Benefits of early detection and treat- ment (based on direct or indirect evidence)	There is no direct evidence on the benefit of screening for HCV infection on health outcomes in asymptomatic adults. There is inadequate direct evidence on the effect of treatment on health outcomes in adults and adolescents. However, there is convincing evidence that the newer DAA regimens result in SVR in a very high proportion (> 95%) of adults aged 18 to 79 years of age and adequate evidence of SVR in adolescents.
	There is adequate evidence of a consistent association between SVR after antiviral therapy and improved health outcomes (decreased risk of all-cause mortality; mortality due to liver disease, cirrhosis, and hepatocellular carcinoma).
	Given the accuracy of the screening test and the availability of effective interventions for HCV infection, the USPSTF determined that the indirect evidence is adequate that the magnitude of the benefit of screening and treatment is substantial for adults aged 18 to 79 years.
Harms of early detection and treatment	Potential harms of screening include anxiety, patient labeling, and feelings of stigmatization. There inadequate direct evidence on the harms of screening for HCV infection.
	Currently recommended DAA regimens are associated with fewer harms than older interferon-containing therapies, and treatment duration is shorter at 8 to 12 weeks. There is adequate evidence that DAA regimens are associated with low rates of serious adverse effects and withdrawal due to adverse effects.
	There is adequate evidence to bound the overall harms of screening and treatment as small based on the known harms of treatment, the high accuracy of screening, and the low likelihood of harms from a blood draw.
USPSTF assessment	The USPSTF concludes with moderate certainty that screening for HCV infection in adults aged 18

Note: For a summary of the evidence systematically reviewed in making this recommendation, the full recommendation statement, and supporting documents, go to https://www.uspreventiveservicestaskforce.org/.

DAA = direct-acting antiviral; HCV = hepatitis C virus; SVR = sustained virologic response; USPSTF = U.S. Preventive Services Task Force.

to 79 years of age has substantial net benefit.

abnormality often includes testing for HCV infection and is considered case finding rather than screening; therefore, it is outside the scope of this recommendation.

SCREENING INTERVALS

Most adults need to be screened only once. Persons with continued risk for HCV infection (e.g., persons who inject drugs) should be screened periodically. There is limited information about the specific screening interval that should occur in persons who continue to be at risk for new HCV infection or how pregnancy changes the need for additional screening.

SCREENING IMPLEMENTATION

Important considerations for implementation of screening include (1) communicating to patients that screening is voluntary and undertaken only with the patient's knowledge and understanding that HCV screening is planned; (2) informing patients about HCV infection, how it can (and cannot) be acquired, the meaning of positive and negative test results, and the benefits and harms of treatment; and (3) providing patients the opportunity to ask questions and to decline screening.

Some health care systems serving insured populations, some academic medical centers, and the Veterans Health Administration have achieved high rates of HCV screening and treatment. However, national HCV screening rates in community health centers and from the National Health Interview Study were 8.3% and 17.3%, respectively; 1 study of 4 safety-net primary care practices serving lowincome and uninsured or underserved populations found that only 0.8% of persons born in 1945 through 1965 were screened over a 1-year period.13 Implementation of successful screening may require addressing various barriers to screening and treatment in diverse populations, such as the uninsured.

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TREATMENT

The purpose of antiviral treatment regimens for HCV infection is to prevent long-term health complications of chronic HCV infection (e.g., cirrhosis, liver failure, hepatocellular carcinoma).

Currently, all oral direct-acting antiviral regimens without interferon have been accepted as the standard treatment for chronic HCV infection. Antiviral therapy is not generally considered during pregnancy because of the lack of data on the safety of newer direct-acting antiviral regimens during pregnancy and breastfeeding. 14,15

ADDITIONAL TOOLS AND RESOURCES

The Centers for Disease Control and Prevention provides strategies for implementing a testing program and additional risk factors at https://www.cdc.gov/hepatitis/hcv/guidelinesc.htm.¹⁶

OTHER RELATED USPSTF RECOMMENDATIONS

The USPSTF has made recommendations on screening for hepatitis B virus infection in pregnant persons,¹⁷ screening for hepatitis B virus infection in adults,¹⁸ and screening for HIV infection.¹⁹

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The "Update of Previous USPSTF Recommendation," "Supporting Evidence," and "Recommendations of Others" sections of this recommendation statement are available at https://www.uspreventiveservicestaskforce.org/uspstf/recommendation/hepatitis-c-screening#fullrecommendationstart.

The USPSTF recommendations are independent of the U.S. government. They do not represent the views of the Agency for Healthcare Research and Quality, the U.S. Department of Health and Human Services, or the U.S. Public Health Service.

References

- 1. Chou R, Dana T, Fu R, et al. Screening for hepatitis C virus infection in adolescents and adults: a systematic review update for the U.S. Preventive Services Task Force. Evidence synthesis no. 188. AHRQ publication 19-05256-EF-1. Agency for Healthcare Research and Quality; 2020.
- Ly KN, Hughes EM, Jiles RB, et al. Rising mortality associated with hepatitis C virus in the United States, 2003-2013. Clin Infect Dis. 2016;62(10): 1287-1288.
- Centers for Disease Control and Prevention. Hepatitis C questions and answers for health professionals. Updated June 18, 2020. Accessed January 23, 2020. https://www.cdc.gov/hepatitis/hcv/hcvfag.htm

- Edlin BR, Eckhardt BJ, Shu MA, et al. Toward a more accurate estimate of the prevalence of hepatitis C in the United States. *Hepatology*. 2015; 62(5):1353-1363.
- Rosenberg ES, Hall EW, Sullivan PS, et al. Estimation of state-level prevalence of hepatitis C virus infection, US States and District of Columbia, 2010. Clin Infect Dis. 2017;64(11):1573-1581.
- Hofmeister MG, Rosenthal EM, Barker LK, et al. Estimating prevalence of hepatitis C virus infection in the United States, 2013-2016. Hepatology. 2019;69(3):1020-1031.
- Centers for Disease Control and Prevention. Surveillance for viral hepatitis—United States, 2017. Updated November 14, 2019. Accessed January 23, 2020. https://www.cdc.gov/hepatitis/ statistics/2017surveillance/index.htm
- 8. U.S. Preventive Services Task Force. Procedure manual. June 2018. Accessed January 23, 2020. https://www.uspreventiveservicestask force.org/Page/Name/procedure-manual
- Chou R, Clark E, Helfand M. Screening for hepatitis C virus infection: systematic evidence review no. 24. Agency for Healthcare Research and Quality; 2004.
- Koneru A, Nelson N, Hariri S, et al. Increased hepatitis C virus (HCV) detection in women of childbearing age and potential risk for vertical transmission—United States and Kentucky, 2011-2014. MMWR Morb Mortal Wkly Rep. 2016;65(28):705-710.
- 11. Ly KN, Jiles RB, Teshale EH, et al. Hepatitis C virus infection among reproductive-aged women and children in the United States, 2006 to 2014. *Ann Intern Med.* 2017;166(11):775-782.
- 12. Chou R, Wasson N. Blood tests to diagnose fibrosis or cirrhosis in patients with chronic hepatitis C virus infection: a systematic review [published correction appears in *Ann Intern Med.* 2013;159(4):308]. *Ann Intern Med.* 2013;158(11):807-820.
- Turner BJ, Rochat A, Lill S, et al. Hepatitis C virus screening and care: complexity of implementation in primary care practices serving disadvantaged populations. Ann Intern Med. 2019;171(12):865-874.
- Tran TT, Ahn J, Reau NS. ACG clinical guideline: liver disease and pregnancy [published correction appears in Am J Gastroenterol. 2016; 111(11):1668]. Am J Gastroenterol. 2016;111(2):176-194, quiz 196.
- 15. American Association for the Study of Liver Diseases, Infectious Diseases Society of America. HCV testing and linkage to care. Updated November 6, 2019. Accessed January 23, 2020. https://www.hcvguidelines.org/full-report/hcv-testing-and-linkage-care
- 16. Centers for Disease Control and Prevention. Testing recommendations for hepatitis C virus infection. Updated April 29, 2020. Accessed January 23, 2020. https://www.cdc.gov/hepatitis/hcv/guidelinesc.htm
- 17. US Preventive Services Task Force. Screening for hepatitis B virus infection in pregnant women: US Preventive Services Task Force reaffirmation recommendation statement [published correction appears in *JAMA*. 2019;322(11):1108]. *JAMA*. 2019;322(4):349-354.
- U.S. Preventive Services Task Force. Screening for hepatitis B virus infection in nonpregnant adolescents and adults: U.S. Preventive Services
 Task Force recommendation statement. Ann Intern Med. 2014;161(1): 58-66.
- US Preventive Services Task Force. Screening for HIV infection: US Preventive Services Task Force recommendation statement. *JAMA*. 2019; 321(23):2326-2336.