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### Title

Risk factors for hepatitis C virus infection at a large urban emergency department

### Permalink

<https://escholarship.org/uc/item/2z44z0g2>

### Journal

Journal of Viral Hepatitis, 29(10)

### ISSN

1352-0504

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### Publication Date

2022-10-01

### DOI

10.1111/jvh.13730

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Peer reviewed

1 **Title: Risk Factors for Hepatitis C Virus Infection at a Large Urban Emergency**

2 **Department**

3

4 **Running Title: Risk Factors for HCV Infection in the ED**

5

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20

21 **Meetings:** None

22

23 **Word Count:** 3,109

24

25 **Author Contributions:** JF, TC, NT and LM conceived the study. LM obtained research  
26 funding. JF and ZM managed and analyzed the data. JF and LM interpreted the data. JF, BS and  
27 EH drafted the manuscript, and all authors contributed substantially to its revision. LM takes  
28 responsibility for the paper as a whole.

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37 **Article Summary:** Independent risk factors for active hepatitis C virus infection in the  
38 emergency department included increasing age, male sex, undomiciled housing status, history of  
39 tobacco use, history of illicit drug use, Medicaid insurance status and Medicare insurance status.

40

41 **Key Words:** HCV, screening, public health, risk factors

- 1 **Data Availability:** The data that support the findings are available from the corresponding
- 2 author upon reasonable request.

**1 ABSTRACT**

2

3 **Background:** In 2020, Centers for Disease Control and Prevention (CDC) released guidelines  
4 recommending HCV screening in all adults 18 years and older. In the current study, we aimed to  
5 identify risk factors for HCV infection in an ED population.

6 **Methods:** We performed a retrospective analysis of ED patients  $\geq 18$  years who were screened  
7 for HCV between November 28, 2018, and November 27, 2019, at a single urban, quaternary  
8 referral academic hospital. An HCV-antibody immunoassay (HCV-Ab) was used for screening;  
9 positive results were confirmed by measuring HCV ribonucleic acid (RNA). The outcome of  
10 interest was the number of new HCV diagnoses (presence of viremia by HCV RNA testing).  
11 Multiple logistic regression models were used to identify risk factors associated with a new HCV  
12 diagnosis.

13 **Results:** 16,722 adult patients were screened for HCV (mean age:  $46 \pm 15$  years; 51% female).  
14 HCV seroprevalence was 5%. Independent risk factors for HCV included increasing age [10-  
15 year aOR 1.26 (95% CI 1.23, 1.30)], male sex [aOR 1.25 (95% CI 1.03, 1.51)], undomiciled  
16 housing status [aOR 2.8 (95% CI 2.3, 3.5)], history of tobacco use [aOR 3.0 (95% CI 2.3, 3.9)],  
17 history of illicit drug use [aOR 3.6 (95% CI 2.9, 4.5)], Medicaid insurance status [aOR 4.0 (95%  
18 CI 2.9, 5.5)] and Medicare insurance status [aOR 1.6 (95% CI 1.1, 2.2)].

19 **Conclusions:** The ED services a high-risk population with regards to HCV infection. These data  
20 support universal screening of ED patients for HCV. Risk factor profiles could improve targeted  
21 screening at institutions without universal testing protocols.

22

# 1 INTRODUCTION

2  
3

## 4 Background

5 There are an estimated 184 million people that have been infected hepatitis C virus (HCV) in the  
6 world<sup>1</sup>. In the United States (U.S.), HCV is the most common and deadliest blood borne  
7 infection in the country, and the number of new acute HCV infections almost tripled between  
8 2011 and 2019 (17,100 to 50,300)<sup>2,3</sup>. As many as 85% of individuals with acute HCV will  
9 become chronically infected, predisposing them to the life-threatening consequences of cirrhosis  
10 and hepatocellular carcinoma<sup>3</sup>. The World Health Organization (WHO) has targeted an 80%  
11 global reduction in new chronic HCV infections by 2030<sup>4</sup>. To meet this goal, population-level  
12 HCV screening initiatives are needed. In 2019, the U.S. Preventative Service Task Force  
13 (USPSTF) released guidelines recommending HCV screening in all adults aged 18-79 years, and  
14 in 2020, the Centers for Disease Control and Prevention (CDC) released guidelines  
15 recommending HCV screening in all adults 18 years and older<sup>5,6</sup>.

16

## 17 Importance

18 The Emergency Department (ED) is an important safety net for underserved, high-risk  
19 populations, making it a vital setting to deliver healthcare services to patients without access to  
20 primary care<sup>7,8</sup>. Compelling evidence continues to emerge on the utility of ED-based infectious  
21 diseases screening programs, including those for Human Immunodeficiency Virus (HIV),  
22 Hepatitis B Virus (HBV), and Hepatitis C Virus (HCV)<sup>8-13</sup>. Indeed, our group recently published  
23 a study that found high HCV antibody (Ab) and HCV RNA seropositivity in ED patients in a  
24 high-volume quaternary care academic center with a large urban and rural catchment area<sup>14</sup>.

1 Additionally, we demonstrated that a universal HCV screening protocol led to a large increase in  
2 HCV testing and new diagnoses<sup>14</sup>. While the results of these studies are encouraging and provide  
3 evidence that the ED setting services a high-risk population, the relatively low prevalence of  
4 HCV in the U.S. (Estimated prevalence: 1%), compared to high-endemic regions such as Central  
5 Asia (Estimated prevalence: 5.8%) and Central Sub-Saharan Africa (Estimated prevalence:  
6 6.0%), makes population-level infection surveillance more difficult and more nuanced<sup>15,16</sup>. There  
7 is evidence to suggest that targeted and birth-cohort based ED-based screening programs may  
8 miss 25% of previously undiagnosed HCV cases, when compared to universal HCV screening<sup>17</sup>.  
9 However, in settings where universal screening programs are unfeasible, tailoring testing  
10 strategies to target high-risk individuals has the potential to improve testing efficiency and reduce  
11 cost. One previous ED-based study examined risk factors for HCV Ab seropositivity, and  
12 another ED-based study examined risk factors for HCV RNA positivity in the 1945-1965 birth  
13 cohort, but to our knowledge, no ED-based studies have examined risk factors for chronic HCV  
14 in a non-targeted adult population<sup>10,11</sup>.

15

## 16 **Goals of This Investigation**

17 In 2018, the study institution implemented an ED-based, universal HCV screening protocol. In  
18 the current study, we aimed to identify risk factors for confirmed HCV infection in an ED  
19 population, with the overall goal of tailoring future screening protocols to the risk profile of the  
20 local community.

21

## 22 **METHODS**

23

## 1 **Study Design and Setting**

### 2 *Overview*

3 The study institution is an academic quaternary referral health system in northern California. The  
4 study ED is a level-1 trauma center in a region with a mixed urban and rural population, and  
5 services more than 80,000 patient visits annually. The study institution implemented an ED-  
6 based HCV screening program on November 27<sup>th</sup>, 2018. We performed a retrospective cohort  
7 analysis of patients who were screened in the ED of the study institution. The overall goal of this  
8 study is to identify risk factors for HCV infection, so that we may better identify high-risk  
9 individuals who are likely to benefit from screening. This study was determined not to be human  
10 subjects research by the study site's Institutional Review Board (IRB) Quality Improvement  
11 Self-Certification Tool.

12

### 13 *Brief Summary of ED-Based HCV Screening Program*

14 All ED patients  $\geq 18$  years who were having blood drawn for any clinical purpose, and who did  
15 not have a positive HCV RNA test result in the electronic medical record (EMR), were eligible  
16 for opt-out HCV screening. Upon entering any blood-based laboratory order into the EMR, a  
17 best practice alert (BPA) notified the ED provider that the patient met screening criteria, at which  
18 point, providers were required to respond to continue with the order entry. If a patient requested  
19 that their insurance not be charged, or they did not have insurance, testing was paid for by the  
20 program grant. Program staff, including two patient navigators (PNs), contacted the patients with  
21 results via telephone or in person, depending on a patient's disposition. Complete details of the  
22 ED-based program have been previously described<sup>18</sup>.

23

## 1 *HCV Laboratory Testing Protocol*

2 An HCV-antibody chemiluminescent immunoassay (HCV-Ab) (Architect i1000, Abbott  
3 Laboratories, Abbott Park, IL) was used to screen patients for HCV. Positive HCV-Ab tests were  
4 confirmed by measuring HCV RNA viral load (Cobas AmpliPrep/TaqMan, Roche Diagnostics,  
5 Basel, Switzerland). HCV-Ab testing typically yielded results within 1-3 days and viral  
6 load testing typically yielded results within 4 days.

7

## 8 **Selection of Participants**

9 We included all adult patients  $\geq 18$  years who were screened for HCV in the ED in the first 12-  
10 months following program implementation (November 28<sup>th</sup>, 2018 to November 27<sup>th</sup>, 2019).

11

## 12 **Measurements**

13 Automated computer-generated reports were used to abstract data directly from the EMR. Data  
14 abstracted included demographic variables, ED visit date, chief complaint, past medical history,  
15 insurance status, and results of HCV testing. Transfusion history was not available. History of  
16 interleukin 28 (IL-28) polymorphism was not available. Each patient was given a unique  
17 identifier to maintain patient confidentiality, and data was stored in de-identified datasets. Only a  
18 patient's first ED visit where they received HCV testing was included in our analysis, to prevent  
19 duplicate data.

20

## 21 **Outcomes**



1 The primary outcomes were the number of new chronic HCV diagnoses (defined as a presence of  
2 viremia by HCV RNA testing) and risk factors for chronic HCV. HCV viremia was used to  
3 define the primary outcome because RNA positive individuals represent the population that can  
4 benefit from linkage-to-treatment. These individuals are also important from a public health  
5 infection control perspective as they are capable of transmitting the virus to others if they are  
6 unaware of their HCV status, and if treated, this can greatly mitigate the transmission of HCV in  
7 the community.

8

## 9 **Analysis**

10 Data were described with simple descriptive statistics. Categorical variables were expressed as  
11 percentages and proportions and continuous variables were expressed as means ( $\pm$  SD). Simple  
12 logistic regression models were used to explore factors trending with new HCV diagnoses  
13 ( $p < 0.1$ ); these factors were then imputed into multiple logistic regression models to assess for  
14 independent association. Factors included in our exploratory analysis included age, sex, race,  
15 ethnicity (Hispanic/non-Hispanic), lesbian-gay-bi-trans-queer (LGBTQ) status, un-domiciled  
16 status, history of HIV, history of tobacco use, history alcohol use, history of illicit drug use and  
17 insurance type (Private/Medicare/Medicaid). Regression outputs were reported as adjusted odds  
18 ratios (aOR) and 95% CIs. Age was analyzed as a continuous variable, and regression outputs  
19 were reported as a ten-year OR. An additional sub-analysis of HCV RNA seroprevalence,  
20 stratified by birth cohort (born 1945-1965, inclusive) and non-birth cohort (born <1945 and  
21 >1965) was performed using the Fisher's Exact test. All analyses were conducted under the  
22 supervision and guidance of a trained biostatistician. Data processing was performed using R

1 4.0.3 (R Core Team, 2020). Statistical analyses were performed using Stata 15.1 (College  
2 Station, TX, USA) and Statistical Analysis Software 9.4M6 (SAS, Institute, Cary, NC).

3

## 4 **RESULTS**

### 5 **Characteristics of Study Subjects**

6 A total of 59,084 adult patients were seen in the ED during the study period, and 46,746 (79%)  
7 underwent phlebotomy for blood-based laboratory studies and were eligible for HCV screening.  
8 A total of 16,722 (36% of eligible patients) adult patients were screened for HCV during the  
9 study period. The mean age of patients was  $46 \pm 15$  years, and most patients were female (51%,  
10  $n=8,578$ ). Race data was available in 99% ( $n=16,531$ ) of patients and ethnicity data was  
11 available in 99.3% ( $n=16,616$ ) of patients. The most common race was White (42%,  $n=6,925$ ),  
12 followed by Mixed/Other (32%,  $n=5,335$ ), Black (20%,  $n=3,267$ ) and Asian (6%,  $n=1,004$ ). The  
13 Hispanic ethnicity comprised 22% ( $n=3,634$ ) of the population. Housing status was known in  
14 88% ( $n=14,754$ ) and eight percent of patients were un-domiciled ( $n=1,184$ ). Three percent of  
15 patients had a known history of HIV (471/16,722). Insurance status was known in all but one  
16 patient. Medicaid insurance was most common (50%,  $n=8,327$ ), followed by Private insurance  
17 (27%,  $n=4,546$ ), Medicare (20%,  $n=3,285$ ), and self-insured/uninsured (3%,  $n=563$ ). Patients  
18 with new diagnoses were older than those with negative HCV testing (mean age:  $54 \pm 12$  years  
19 vs.  $46 \pm 16$ ). Most new HCV diagnoses were male (66%), White (50%), undomiciled (30%), and  
20 had Medicaid insurance (68%). Full patient characteristics, stratified by HCV infection status,  
21 are described in **Table 1**.

22

## 1 HCV Testing Results

2 Nine percent (1,519/16,722, 95% CI 8.7, 9.5) of patients tested positive for HCV-Ab. All  
3 patients with positive HCV-Ab testing received reflex HCV RNA testing. Confirmatory HCV  
4 RNA testing was reactive in 54% (814/1,519, 95% CI 51, 56) of patients who initially screened  
5 positive for HCV, yielding 814 new HCV diagnoses (overall prevalence: 5%). The  
6 seroprevalence of confirmed HCV was 6.6% in males and 3.3% in females. The seroprevalence  
7 of confirmed HCV by race was 5.9% in Whites, 5.5% in blacks, 4% in Other/mixed and 1% in  
8 Asians. A table with full HCV Ab and RNA seroprevalence data in sub-groups, is available in  
9 the **Supplemental Table**.

10

## 11 HCV Risk Factor Analysis

12 In our simple logistic regression model age, sex, race, ethnicity, housing status, history of  
13 tobacco use, history of alcohol use, history of illicit drug use and insurance type all met the *a*  
14 *priori* cutoff level for significance ( $p < 0.1$ ) and were included in our multivariate model (Table 2).  
15 In our multiple logistic regression model, independent risk factors for an HCV positive status  
16 included increasing age [10-year aOR 1.26 (95% CI 1.23, 1.30)], male sex [aOR 1.25 (95% CI  
17 1.03, 1.51)], undomiciled housing status [aOR 2.8 (95% CI 2.3, 3.5)], history of tobacco use  
18 [aOR 3.0 (95% CI 2.3, 3.9)], history of illicit drug use [aOR 3.6 (95% CI 2.9, 4.5)], Medicaid  
19 insurance status [aOR 4.0 (95% CI 2.9, 5.5)] and Medicare insurance status [aOR 1.6 (95% CI  
20 1.1, 2.2)]. Relative to the white race, the Black and Asian races were independent protective  
21 factors for an HCV positive status [Black race, aOR 0.78 (95% CI 0.62, 0.98); Asian race, aOR  
22 0.3 (0.1, 0.6)].

1

## 2 **Birth Cohort Sub-Analysis**

3 HCV diagnoses increased with each decade of age, peaking at 58-67 years, before declining  
4 again (Figure 1). HCV diagnoses were more common in the 1945-1965 birth cohort compared to  
5 the non-birth cohort (7.8% vs. 3%,  $p < 0.001$ ).

6

## 7 **DISCUSSION**

8

9 To our knowledge, the current study provides the largest cohort of ED-based HCV screening to  
10 date. In the US, an estimated 1.7% of individuals are HCV-Ab positive and 1% of individuals  
11 have confirmed HCV<sup>15</sup>. In the current study, 9% of individuals tested positive for HCV-Ab, and  
12 5% of individuals had RNA-confirmed HCV, which is similar to that reported by other large,  
13 non-targeted, ED-based HCV screening studies (HCV-Ab+: 6-13.2%; HCV RNA+: 1.2-7.7%)<sup>11-  
14 13,19</sup>. These data suggest that, compared to the general population, the ED services a high-risk  
15 population with regards to HCV exposure and seroconversion.

16

17 In 2020, the CDC updated its previous HCV screening guidelines from targeted screening in the  
18 birth cohort 1945-1965, to universal screening of all adults 18 years and older<sup>5,20</sup>. In our study,  
19 the 1945-1965 birth cohort (which corresponds to an approximate age of 53-73 in our study) had  
20 a much higher rate of confirmed HCV, than those who were born either before 1945 or after  
21 1965. It is estimated that 2.6% of birth-cohort patients in the US have chronic HCV<sup>20</sup>. In one  
22 recent ED-based study of an HCV birth-cohort screening program, approximately 2.3% of  
23 patients had confirmed HCV, whereas in our birth cohort sub-analysis, 7.8% of patients had

1 RNA-confirmed HCV<sup>10</sup>. Given the results of our birth-cohort sub-analysis, it was not surprising  
2 that increasing age was an independent predictor of HCV positivity. However, it was notable that  
3 the prevalence of confirmed HCV was high even in patients born after 1965 (3.1% for patients  
4 28-47 years), which was 3-fold higher than the overall US HCV prevalence (~1%)<sup>15</sup>. These data  
5 support the current CDC recommendation to screen non-birth cohort individuals.

6

7 Although increasing age was associated with an increased risk of HCV, peaking in the 58-67  
8 year-old sub-group, the prevalence fell in the 68-77 year-old sub-group. Thus, it was surprising  
9 that Medicare insurance was an independent risk factor for HCV positivity. The estimated  
10 prevalence of HCV in the Medicare population ranges between 1.2-3.2%, compared to 5.4% in  
11 our study<sup>21,22</sup>. In the Medicare population, HCV is associated with increased all-cause mortality  
12 and increased healthcare resource utilization<sup>21,23</sup>. As the baby boomer generation ages,  
13 downstream clinical sequelae of HCV are likely to have substantive epidemiologic and economic  
14 repercussions, highlighting the need for continued screening and treatment initiatives within this  
15 age group.

16

17 US national data demonstrates that Medicaid patients have the highest prevalence of HCV  
18 (2.6%), compared to other insurance groups (private, self-insured, Medicare etc.)<sup>22</sup>. In our study,  
19 patients with a Medicaid insurance status had a prevalence of 7.2%, which was nearly 2.5-fold  
20 higher than those with private insurance. Unsurprisingly, Medicaid status, a crude approximation  
21 for low socioeconomic status (SES) was independently associated with having confirmed HCV.  
22 These data are particularly concerning, given that adjusted survival analyses demonstrate that  
23 among patients with HCV, those with Medicaid have much higher mortality<sup>22</sup>. Given the

1 accessibility of curative treatment and the high-risk nature of HCV in this population, more  
2 robust screening efforts must be established in order to connect these patients to appropriate  
3 outpatient care.

4  
5 In pooled national data, males account for the majority of HCV infections in the U.S., with a  
6 male to female HCV prevalence ratio of 2.3 (prevalence: males: 1.31%, females: 0.57%)<sup>24</sup>. In our  
7 study, the male sex had an HCV prevalence over twice that of the female sex (6.6% vs. 3.2%),  
8 and was an independent risk factor for confirmed HCV infection. This is consistent with one  
9 previous ED-based study that explored HCV risk factors in a 1945-1965 birth cohort<sup>10</sup>. The  
10 Black race has been previously identified as a risk factor for HCV, with national data  
11 demonstrating a prevalence that is 2-3 fold times higher than other racial groups<sup>10,24</sup>. A recent  
12 study of four urban emergency departments showed that anti-HCV positivity was higher in  
13 Whites than Blacks, although the analysis did not adjust for other important co-variates, so it is  
14 unclear if this association was linked to race alone<sup>25</sup>. In our study, HCV prevalence was similar  
15 between Whites (5.9%) and Blacks (5.5%). Interestingly, compared to the white race in our  
16 multivariate risk analysis, the black race actually appeared to be predictive of a decreased risk of  
17 HCV. The Asian race was also independently associated with a decreased risk of HCV and had  
18 the lowest seroprevalence of anti-HCV antibody (1.9%) and HCV RNA (1%) among all racial  
19 sub-groups. While published HCV data on Asian-American populations are sparse, in one  
20 community-based study of Asian-Americans in California, anti-HCV-Ab seroprevalence was  
21 5.5%, compared to 2.3% for non-Asians (HCV RNA seroprevalence was not reported)<sup>26</sup>. There  
22 are many possible explanations for the discrepancy between previously published reports and our  
23 data, but one possibility is that Asian-Americans seem to make up a disproportionate minority of

1 ED visits (5%), relative to the proportion of Asian-Americans living in the local county (17%),  
2 resulting in a sampling bias<sup>27</sup>. Alternatively, local HCV prevalence of Asian Americans may be  
3 influenced by the ethnic makeup of this subpopulation, as evidence exists that HCV  
4 seroprevalence differs drastically by country of origin, and is particularly high in those of  
5 Vietnamese heritage, which makes up just 10% of the Asian-American population in the local  
6 county<sup>26,28,29</sup>. Alternatively, there may be unexplored lifestyle or socioeconomic factors within  
7 this racial group that are confounding this finding. Future studies should explore race as an effect  
8 modifier for HCV risk factors in the ED setting.

9  
10 In 2020, over half a million Americans were undomiciled, a figure that is expected to rise over  
11 the coming decade<sup>30</sup>. California comprises 12% of the U.S. population but accounts for 27% of  
12 the nation's homeless population<sup>31</sup>. One previous ED-based study found that homelessness was a  
13 risk factor for HCV-Ab seropositivity, but did not explore its relationship with confirmed HCV  
14 (detectable HCV RNA)<sup>11</sup>. The seroprevalence of anti-HCV Ab in the homeless population in the  
15 US ranges from 19-69%, but few studies report viral-load confirmed HCV<sup>32</sup>. However, one study  
16 of five health centers that serviced predominantly homeless populations, found a confirmed HCV  
17 prevalence of 8.4% in this population, which was over 50% less than the HCV prevalence  
18 reported in our undomiciled sub-group (18.9%)<sup>33</sup>. In our study, an undomiciled status was an  
19 independent risk factor for RNA-confirmed HCV, and undomiciled individuals were almost 5-  
20 fold more likely to have a new HCV diagnosis compared to domiciled individuals. Homelessness  
21 has a well-documented two-way association with illicit and injection drug use, with some studies  
22 suggesting that homelessness is a risk factor for future injection drug use, and others suggesting  
23 that injection drug use increases the likelihood of future homelessness<sup>34-37</sup>. Interestingly, both an

1 undomiciled status and history of illicit drug use were independently associated with HCV  
2 infection, suggesting that there may additional unmeasured behavioral factors in the undomiciled  
3 population. A US-based study found that a history of illicit drug use (including IV drug use) was  
4 independently associated with chronic HCV, and the prevalence of HCV in this cohort was  
5 5.6%<sup>38</sup>. A cross-sectional study in the ED found that patients who injected drugs were almost 16-  
6 fold time likely to have HCV<sup>39</sup>. In our study, the prevalence of confirmed HCV in those who  
7 used illicit drugs was 10.4%, indicating that a substance use epidemic may be contributing to  
8 rising local HCV infections.

9  
10 While ED-based, non-targeted HCV screening programs have been shown to be successful in  
11 both academic and community settings, many emergency departments may not the resources  
12 necessary to conduct universal screening<sup>14,40</sup>. However, targeted screening interventions are  
13 likely to be feasible even in EDs with fewer resources. Thus, we believe that our methodology  
14 may be used as a model for developing local HCV risk factor profiles, which can be used to  
15 target high-risk ED patients in settings without universal screening protocols.

16  
17 Our study must be interpreted in light of its limitations. This was a single-center study at a large,  
18 academic center with a mixed urban and rural population, thus, our findings may not be  
19 generalizable to all settings. This study is limited by its retrospective design, which affected the  
20 completeness of certain variables. Finally, data related to certain well-documented HCV risk  
21 factors, such as blood transfusion history and sexual history, were not available and were not  
22 explored in this ED-based analysis<sup>5</sup>. IL-28 polymorphisms have been associated with  
23 spontaneous clearance of HCV; we did not have data on the presence of this polymorphism, and



1 thus could not include it in our model<sup>41</sup>. We do not explore long-term, patient-centered outcomes  
2 in this study. Future studies could examine an estimation in the gain in quality-adjusted life years  
3 (QALYs) from early HCV diagnosis in the ED setting.

4

5 In summary, the study institution's ED services a high-risk population with regards to HCV  
6 positivity. Increasing age, male sex, undomiciled housing status, history of tobacco use, history  
7 of illicit drug use, and Medicaid and Medicare insurance were all independent risk factors for  
8 HCV positivity. The Black and Asian races had a lower risk of HCV, compared to the White  
9 race. Overall, HCV seroprevalence was high in all sub-groups, which supports universal testing  
10 in this population. However, risk factor profiles could improve targeted screening at institutions  
11 without universal HCV testing protocols.

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1 **CONFLICTS OF INTEREST**

2

3 **Conflict of Interest Disclosure:** The authors have no conflicts of interest to disclose.

4

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2 **ACKNOWLEDGEMENTS:**

3

4 **Funding Support:** This work was supported by Gilead’s FOCUS program. In the U.S., the  
5 FOCUS Program is a public health initiative that enables partners to develop and share best  
6 practices in routine blood-borne virus (HIV, HCV, HBV) screening, diagnosis, and linkage to  
7 care in accordance with screening guidelines promulgated by the U.S. Centers for Disease  
8 Control and Prevention (CDC), the U.S. Preventive Services Task Force (USPSTF), and state  
9 and local public health departments. FOCUS funding supports HIV, HCV, and HBV screening  
10 and linkage to a first appointment. FOCUS partners do not use FOCUS awards for activities  
11 beyond linkage to a first appointment.

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1 **Figure Legends**

2 **Figure 1.** A. Hepatitis C Virus (HCV) antibody (Ab) and RNA viral load (VL) seroprevalence  
3 by age group. B. HCV Ab and HCV RNA VL seroprevalence by birth cohort (BC, born 1945-  
4 1965) and non-BC cohort (born <1945 and >1965).

**Table 1.** Patient characteristics stratified by HCV infection status

<b>Characteristic</b>	<b>New HCV Diagnosis<sup>1</sup> (n=814)</b>	<b>HCV Negative (n=15,908)</b>	<b>P</b>
<b>Age (years)<sup>2</sup></b>	54 ( $\pm$ 12)	46 ( $\pm$ 16)	<0.001
<b>Male sex</b>	66% (541/814)	48% (7,601/15,906)	<0.001
<b>Race</b>			
<i>White</i>	50% (407/809)	41% (6,518/15,722)	<0.001
<i>African American</i>	22% (179/809)	20% (3,088/15,722)	0.4
<i>Asian</i>	1% (10/809)	6% (994/15,722)	<0.001
<i>Mixed/Other</i>	26% (213/809)	33% (5,122/15,722)	<0.001
<b>Hispanic Ethnicity</b>	17% (138/810)	22% (3,496/15,752)	<0.001
<b>LGBTQ</b>	4% (2/52)	5% (177/3,428)	1.0
<b>Un-domiciled Status</b>	30% (221/741)	7% (963/14,013)	<0.001
<b>History of HIV</b>	3% (26/814)	3% (445/15,908)	0.5
<b>History of Tobacco Use</b>	76% (608/799)	42% (6,554/15,462)	<0.001
<b>History of Alcohol Use</b>	65% (431/667)	53% (6,666/12,614)	<0.001
<b>History of Illicit Drug Use</b>	73% (499/681)	34% (4,277/12,639)	<0.001
<b>Insurance Type</b>			
Private	8% (68/814)	28% (4,478 /15,907)	<0.001
Medicare	22% (176/814)	20% (3,109/15,907)	0.001
Medicaid	68% (551/814)	49% (7,776/15,907)	<0.001
Self/Uninsured	2% (19/814)	3% (544/15,907)	0.001

<sup>1</sup> Defined as positive HCV antibody *and* HCV viral load.

<sup>2</sup> Reported as mean  $\pm$  standard deviation

HCV, hepatitis C virus; HIV, human immunodeficiency virus; LGBTQ, lesbian-gay-bisexual-trans-queer;

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**Table 2.** Simple and multiple logistic regression models identifying risk factors for new HCV diagnoses

Predictive Factor	Bivariate Model		Multivariate Model	
	aOR (95% CI)	P	aOR (95% CI)	P
<b>Age (10 years)<sup>1</sup></b>	1.02 (1.01, 1.02)	<0.001	1.26 (1.23, 1.30)	<0.001
<b>Male sex</b>	2.2 (1.9, 2.5)	<0.001	1.25 (1.03, 1.51)	0.02
<b>Race</b>				
<i>White (Ref)</i>	-	-		
<i>Black</i>	0.9 (0.8, 1.1)	0.4	0.78 (0.62, 0.98)	0.03
<i>Asian</i>	0.2 (0.1, 0.3)	<0.001	0.3 (0.1, 0.6)	0.002
<i>Mixed/Other</i>	0.7 (0.6, 0.8)	<0.001	0.9 (0.7, 1.2)	0.4
<b>Hispanic</b>	0.7 (0.6, 0.9)	0.001	1.0 (0.8, 1.4)	0.9
<b>Undomiled Status</b>	5.8 (4.9, 6.8)	<0.001	2.8 (2.3, 3.5)	<0.001
<b>History of HIV</b>	1.1 (0.8, 1.7)	0.5	-	-
<b>History of Tobacco Use</b>	4.3 (3.7, 5.1)	<0.001	3.0 (2.3, 3.9)	<0.001
<b>History of Alcohol Use</b>	1.6 (1.4, 1.9)	<0.001	0.9 (0.8, 1.1)	0.4
<b>History of Illicit Drug Use</b>	5.4 (4.5, 6.4)	<0.001	3.6 (2.9, 4.5)	<0.001
<b>Insurance Type</b>				
<i>Private (Ref)</i>	-	-	-	-
<i>Medicare</i>	3.7 (2.8, 5.0)	<0.001	1.6 (1.1, 2.2)	0.01
<i>Medicaid</i>	4.7 (3.6, 6.0)	<0.001	4.0 (2.9, 5.5)	<0.001
<i>Self/Uninsured</i>	2.3 (1.4, 3.9)	0.002	2.0 (0.9, 4.5)	0.08

Factors trending with the outcome of interest (new HCV diagnosis) in the simple logistic regression (p<0.1) were included in the multiple logistic regression model to assess for independent association.

<sup>1</sup>Age analyzed as continuous variable with output reported as 10-year ORs

aOR, adjusted odds ratio. HIV, human immunodeficiency virus. Ref, reference variable.