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Original article

The impact of hepatitis C on labor force participation, absenteeism, presenteeism and non-work activities

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Abstract

Objective:

Between 2.7 and 3.9 million people are currently infected with the hepatitis C virus (HCV) in the United States. Although many studies have investigated the impact of HCV on direct healthcare costs, few studies have estimated the indirect costs associated with the virus using a nationally-representative dataset.

Methods:

Using data from the 2009 United States (US) National Health and Wellness Survey, patients who reported a hepatitis C diagnosis ($n = 695$) were compared to controls on labor force participation, productivity loss, and activity impairment after adjusting for demographics, health risk behaviors, and comorbidities. All analyses applied sampling weights to project to the population.

Results:

Patients with HCV were significantly less likely to be in the labor force than controls and reported significantly higher levels of absenteeism (4.88 vs. 3.03%), presenteeism (16.69 vs. 13.50%), overall work impairment (19.40 vs. 15.35%), and activity impairment (25.01 vs. 21.78%). A propensity score matching methodology replicated many of these findings.

Conclusions:

While much of the work on HCV has focused on direct costs, our results suggest indirect costs should not be ignored when quantifying the societal burden of HCV. To our knowledge, this is the first study which has utilized a large, nationally-representative data source for identifying the impact of HCV on labor force participation and work and activity impairment using both a propensity-score matching and a regression modeling framework.

Limitations:

All data were patient-reported (including HCV diagnosis and work productivity), which could have introduced some subjective biases.

Introduction

The hepatitis C virus (HCV) is a blood-borne illness which typically manifests as fever, fatigue, abdominal pain, and jaundice (among other symptoms), though as many as 80% of the newly infected are asymptomatic¹. The incidence of HCV has declined in recent decades, with the current rate estimated at 0.3 cases per 100,000². It is believed that between 2.7 and 3.9 million people are currently infected with HCV in the United States³. In the later stages of infection, patients with HCV are at increased risk for the development of cirrhosis and

hepatocellular carcinoma which are associated with high morbidity and mortality rates⁴. HCV is the primary contributor to deaths from liver disease⁵.

A number of studies have evaluated the impact of HCV on healthcare costs^{6–8}. Previous studies have estimated direct healthcare costs associated with HCV to be \$2,070 per patient in 1997⁶ and \$2,470 per patient during the time period from 1997–1999⁸. However, direct medical costs represent only part of the societal burden of HCV infection. Indirect costs, the costs attributed to work impairment (either through health-related absences or health-related impairment while working), are also important to consider, but have been largely ignored in the HCV literature.

Previous cost-effectiveness models have left out work impairment altogether⁹ or have measured productivity losses only in the form of premature mortality and disability as a result of projected late stage liver disease⁷. One of the only studies to analyze current (rather than projected) costs related to HCV found total indirect costs of \$3.66 billion in 1997, though the algorithms used focused exclusively on wages, fringe benefits, and home production⁶. A review of the literature yielded only one study each that assessed the impact of HCV on labor force participation¹⁰, employability in a welfare population¹¹, and the impact of HCV on absenteeism and presenteeism¹². Apart from these studies, the only assessments of HCV impact on workplace activity have been within the context of clinical trials¹³.

To fully understand the societal impact of HCV, the association between the virus and both labor force participation and work productivity loss must be considered. It is also crucial to investigate potential confounding variables that may contribute to a relationship between HCV status and workplace activity. A variety of studies have documented the impact comorbidities and health behaviors may have on health outcomes among HCV patients, including psychiatric illnesses^{14–16}, advanced fibrosis¹⁷, fatigue¹⁸, and depressive symptoms^{19,20}. As these variables all may plausibly affect labor force participation and work productivity, they should be considered as potential confounders. As such, the present study attempts to determine the incremental effect of HCV on labor force participation, work productivity loss, and activity impairment using a large, nationally-representative database, and accounting for many of the differences between the groups that were previously ignored. These results will help provide a robust estimate of the contribution of HCV to indirect costs.

Methods

National Health and Wellness Survey

The current study used data from the 2009 wave ($N = 75,000$) of the US National Health and Wellness

Survey (NHWS; Kantar Health, New York, NY, USA), an annual, cross-sectional study of adults aged 18 years or older. The NHWS includes epidemiological data, treatment information, information on health risk behaviors, and health-related outcome data. Potential respondents to NHWS are recruited through an existing consumer panel. The consumer panel recruits its panel members through opt-in emails, co-registration with panel partners, e-newsletter campaigns, and banner placements. All panelists must explicitly agree to be a panel member, register with the panel through a unique email address, and complete an in-depth demographic registration profile. Using a stratified random sample framework (with quotas based on gender, age, and race/ethnicity), the demographic composition of the 2009 US NHWS sample is comparable to that of the US adult population as described by the March 2008 Current Population Survey of the US Census Bureau²¹. Additional comparisons with NHWS and NHIS have been made elsewhere²². All subjects provided informed consent and the study was approved by Essex Institutional Review Board (Lebanon, NJ, USA). In the NHWS study, 501,239 subjects were contacted, out of whom 92,759 responded (18.5% response rate). Of those who responded, 75,000 patients gave informed consent, met inclusion criteria, and completed the survey. Patients diagnosed with hepatitis B, HIV, or AIDS ($n = 966$) were excluded from the sample, leaving a total sample size of 74,034. Because the base rates of these conditions are low among the general population yet relatively high among the HCV population, patients diagnosed with these conditions were excluded to ensure any observed differences between HCV diagnosed patients and controls were not due to HCV-related comorbidities. This has also been done in prior research²¹. Of the total 74,034 respondents, 695 reported being diagnosed with HCV and 73,339 reported not being diagnosed with HCV.

Labor force and work productivity variables

The potential impact of HCV was assessed in terms of the following five workplace and associated characteristics: (1) labor force participation (whether or not the respondent is employed/actively seeking work); (2) absenteeism (percentage of work time missed due to health); (3) presenteeism (degree of impairment experienced at work due to health); (4) overall work impairment (a total percentage of missed work time due to either absenteeism or presenteeism); and (5) activity impairment (degree of impairment experienced during non-work activities).

Labor force participation

All respondents reported their current workforce status. Those who reported being employed full-time, employed part-time, self-employed, or unemployed but looking for

work were considered to be in the labor force. All others were not considered to be in the labor force.

Work productivity and activity impairment

The Work Productivity and Activity Impairment (WPAI) questionnaire was used to measure the impact of health on employment-related activities²³. WPAI is a 6-item validated instrument that consists of four metrics: absenteeism (the percentage of work time missed because of one's health in the past 7 days), presenteeism (the percentage of impairment experienced while at work in the past 7 days because of one's health), overall work productivity loss (an overall impairment estimate that is a combination of absenteeism and presenteeism), and activity impairment (the percentage of impairment in daily activities because of one's health in the past 7 days). Only respondents who reported being employed full-time, employed part-time, or self-employed provided data for absenteeism, presenteeism, and overall work impairment. All respondents provided data for activity impairment. The validity and accuracy of the instrument has been established in a number of disease states, included HCV¹³.

Absenteeism was calculated by dividing the number of work hours a patient missed in the past week because of his or her health by the total number of hours a patient could have worked (the number of hours he/she did work plus the number of hours missed because of his/her health) and converting this proportion into a percentage. Presenteeism was measured by a patient's rating of his or her level of impairment experienced while at work in the past 7 days (from 0 to 10, with higher numbers indicating greater impairment), which was then multiplied by 10 to create a percentage, with a range from 0% to 100%. Overall work impairment was measured by adding absenteeism and presenteeism to determine the total percentage of lost work time. Activity impairment was measured by a patient's rating of the level of impairment experienced in daily activities in the past 7 days (from 0 to 10, with higher numbers indicating greater impairment), which was then multiplied by 10 to create a percentage, with a range from 0 to 100%.

Control variables

A wide array of predictor variables that have previously been shown to have a potential impact on both labor force participation as well work productivity loss was evaluated. These variables were grouped into socio-demographic variables, health-risk behaviors, and morbidity/comorbidity status variables. The socio-demographic variables included gender, race/ethnicity (white, black, Hispanic, Asian, or other), marital status (married/living with partner vs. all else), educational attainment (college degree vs. all else), and annual household income (<\$25,000, \$25,000 to

<\$50,000, \$50,000 to <\$75,000, \$75,000 or more, decline to answer). Health-risk behaviors included tobacco smoking, alcohol consumption, body mass index (BMI) and physical exercise. For the assessment of morbidity/comorbidity status, both the number of comorbid conditions and, separately, the self-reported presence of anxiety and depression were assessed.

Statistical analysis

Initial bivariate comparisons between HCV and controls were made using chi-square tests for categorical outcomes and independent-samples *t*-tests for continuous outcomes. Because some of the outcomes were available for all respondents (labor force participation and activity impairment) and some outcomes were only available for those who were employed (absenteeism, presenteeism, and overall work impairment), different analysis groups were created (see Figure 1). The primary analytical approach was regression modeling. The regression modeling approach included patients diagnosed with HCV and controls. Age, gender, race/ethnicity, education, income, health insurance, presence of anxiety, presence of depression, number of comorbidities, smoking status, exercise behavior, alcohol use, BMI and group membership (HCV group vs. control group) were included in a series of generalized linear models (specifying a negative binomial distribution and a log-link function) to predict work productivity and activity impairment metrics. A log-link function was selected because of the pronounced non-normality of the dependent variables (high positive skew) and the negative binomial distribution was selected because it best modeled the dispersion. A thorough discussion of these approaches is made elsewhere²⁴. A logistic regression model (with the same independent variables) was used to predict labor force participation (yes vs. no). The formula for these models is as follows (with a log or logit depending upon the outcome):

$$\begin{aligned} &\log(Y_{\text{Work productivity}}) \text{ or } \text{logit}(Y_{\text{Labor force}}) \\ &= b_0 + b_{\text{Age}}(X_{\text{Age}}) + b_{\text{Female gender}}(X_{\text{Female gender}}) \\ &\quad + b_{\text{Black ethnicity}}(X_{\text{Black ethnicity}}) + b_{\text{Hispanic ethnicity}} \\ &\quad \times (X_{\text{Hispanic ethnicity}}) + b_{\text{Asian ethnicity}}(X_{\text{Asian ethnicity}}) \\ &\quad + b_{\text{Other ethnicity}}(X_{\text{Other ethnicity}}) + b_{\text{Married}}(X_{\text{Married}}) \\ &\quad + b_{\text{College educated}}(X_{\text{College educated}}) + b_{\text{Income} < \$25K} \\ &\quad \times (X_{\text{Income} < \$25K}) + b_{\text{Income} \$50K \text{ to } < \$75K} \\ &\quad \times (X_{\text{Income} \$50K \text{ to } < \$75K}) + b_{\text{Income} \$75K \text{ or more}} \\ &\quad \times (X_{\text{Income} \$75K \text{ or more}}) + b_{\text{Decline to answer income}} \\ &\quad \times (X_{\text{Decline to answer income}}) + b_{\text{Health insurance}}(X_{\text{Health insurance}}) \\ &\quad + b_{\text{Anxiety}}(X_{\text{Anxiety}}) + b_{\text{Depression}}(X_{\text{Depression}}) \end{aligned}$$

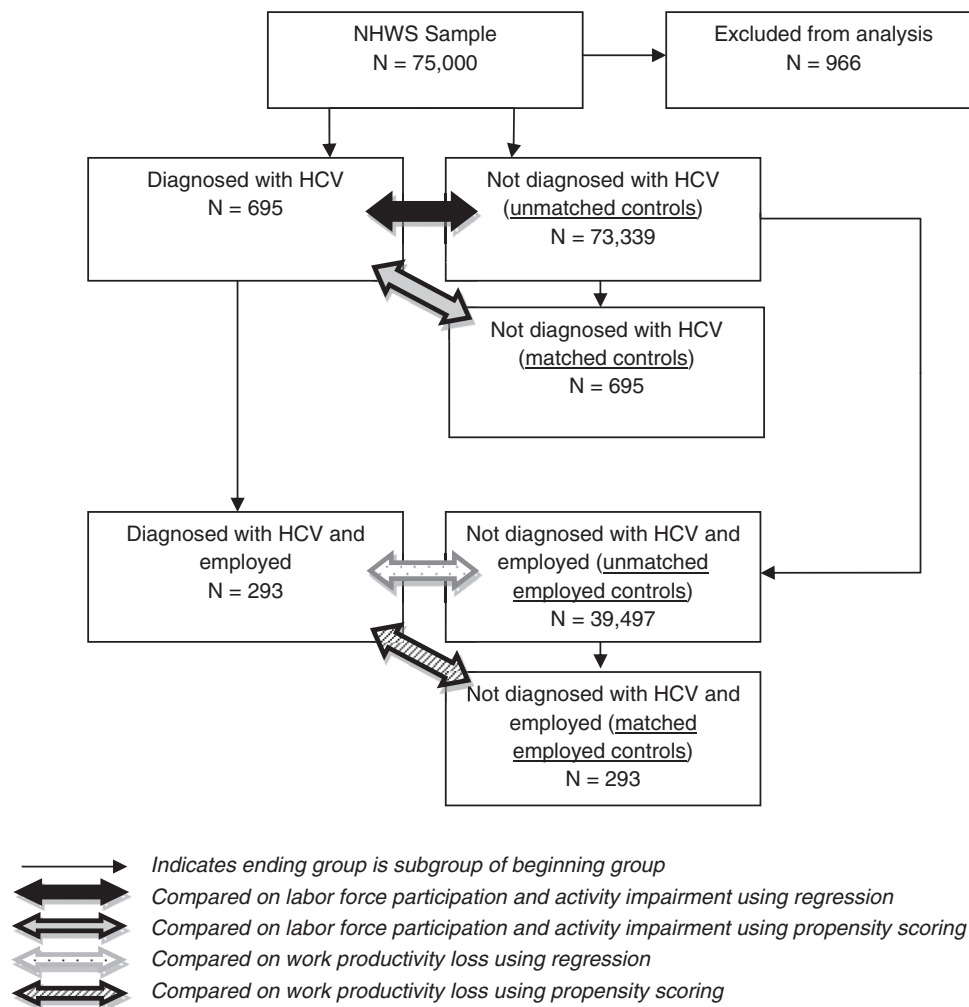


Figure 1. Overview of analysis groups and methods.

$$\begin{aligned}
 &+ b_{\text{Comorbidities}}(X_{\text{Comorbidities}}) + b_{\text{Smoking status}}(X_{\text{Smoking status}}) \\
 &+ b_{\text{Exercise behavior}}(X_{\text{Exercise behavior}}) + b_{\text{Alcohol use}} \\
 &\times (X_{\text{Alcohol use}}) + b_{\text{BMI:underweight}}(X_{\text{BMI:underweight}}) \\
 &+ b_{\text{BMI:overweight}}(X_{\text{BMI:overweight}}) + b_{\text{BMI:obese}}(X_{\text{BMI:obese}}) \\
 &+ b_{\text{BMI:decline to answer}}(X_{\text{BMI:decline to answer}}) \\
 &+ b_{\text{HCV status}}(X_{\text{HCV status}}) + \text{error}
 \end{aligned}$$

Regression techniques used in situations such as this do present some disadvantages. The analysis groups are of vastly different sizes and the characteristics of the HCV group may be so meaningfully different that the extrapolation conducted during the regression process may not be an accurate estimate of the effect of HCV. To combat these limitations, and provide a more robust analysis of HCV, a propensity score modeling approach was also used to replicate the findings.

Two separate propensity score matching procedures were used. In the first matching procedure, all HCV

patients were matched to a subset of all controls. Age, gender, race/ethnicity, education, income, health insurance, presence of anxiety, presence of depression, number of comorbidities, smoking status, exercise behavior, alcohol use, and BMI were entered in a logistic regression to predict group membership (HCV group vs. unmatched control group) in order to obtain propensity score values. Next, each HCV patient was matched with the control patient with the closest propensity score using a greedy-matching algorithm²⁵. Post-match, these groups were compared on labor force participation and activity impairment (the two outcomes that were answered by all respondents).

In the second matching procedure, employed HCV patients were matched to a subset of employed controls. The same variables noted above were entered in a logistic regression to predict group membership (employed HCV group vs. unmatched employed control group) in order to obtain propensity score values. Next, each employed HCV

Table 1. Demographic and health history differences of hepatitis C patients compared to both unmatched and matched controls.

Variable	Hepatitis C group <i>n</i> = 695				Unmatched control group <i>n</i> = 73,339				<i>p</i> -value
	Unweighted <i>n</i>	Weighted			Unweighted <i>n</i>	Weighted			
		<i>n</i>	%	SE (%)		<i>n</i>	%	SE (%)	
Female	286	865,034	41.9	±1.98	37,964	114,086,430	52.2	±0.21	<0.0001
White	510	1,390,781	67.4	±2.03	54,412	150,917,684	69.0	±0.21	0.449
Black	76	223,617	10.8	±1.20	7,494	24,668,483	11.3	±0.14	0.7198
Hispanic	60	306,387	14.9	±1.84	5,947	28,768,182	13.2	±0.16	0.3604
Asian	6	16,465	0.8	±0.33	3,430	8,448,284	3.9	±0.08	<0.0001
Other race	43	125,310	6.1	±0.91	2,056	5,962,965	2.7	±0.08	0.0003
Married/living with partner	383	1,124,619	54.5	±2.02	44,978	129,990,087	59.4	±0.20	0.0169
College educated	166	466,643	22.6	±1.65	26,887	78,897,185	36.1	±0.20	<0.0001
Employed	293	882,674	42.8	±1.99	39,497	122,562,298	56.0	±0.21	<0.0001
Have health insurance	536	1,582,740	76.7	±1.67	60,466	176,869,058	80.8	±0.16	0.0145
Income <\$25 k	225	655,725	31.8	±1.84	13,299	41,247,829	18.9	±0.17	<0.0001
Income \$25 k to <\$50 k	224	672,893	32.6	±1.89	21,952	65,443,921	29.9	±0.19	0.1543
Income \$50 k to <\$75 k	123	373,812	18.1	±1.67	15,749	46,376,502	21.2	±0.16	0.0665
Income \$75 k+	100	294,901	14.3	±1.39	17,884	52,614,301	24.1	±0.17	<0.0001
Decline to answer	23	65,228	3.2	±0.67	4,455	13,083,045	6.0	±0.10	<0.0001
Currently smoke	358	1,114,186	54.0	±2.00	16,286	49,236,764	22.5	±0.17	<0.0001
Drink alcohol	379	1,141,257	55.3	±2.00	47,698	143,079,030	65.4	±0.20	<0.0001
Currently exercise	357	1,061,104	51.4	±2.02	46,547	140,889,633	64.4	±0.20	<0.0001
Comorbid anxiety	251	756,353	36.7	±1.92	11,451	35,110,320	16.0	±0.15	<0.0001
Comorbid depression	294	885,238	42.9	±1.99	12,907	38,960,073	17.8	±0.15	<0.0001
	Mean	Mean	SD	SE	Mean	Mean	SD	SE	<i>p</i> -value
Age	51.46	50.35	11.02	0.42	47.89	45.96	16.93	0.06	<0.0001
Body mass index (BMI)	29.71	29.78	7.45	0.28	28.78	28.62	7.04	0.03	<0.0001
Number of comorbidities	8.14	8.25	6.29	0.24	4.34	4.18	4.49	0.02	<0.0001

Unweighted refers to statistics which did not apply a sampling weight; weighted refers to statistics which applied a sampling weight.
SE, standard error; SD, standard deviation.

patient was matched with the employed control patient with the closest propensity score using a greedy-matching algorithm²⁵. Post-match, these groups were compared on absenteeism, presenteeism, and overall work impairment (the three outcomes that were answered by only those who were employed).

Generalized linear models, specifying a negative binomial distribution and a log-link function, were used to analyze group differences on work and activity impairment variables because of the pronounced non-normality of the data. Since employed HCV patients differed from matched employed controls on prevalence of smoking, this variable was controlled for when predicting absenteeism, presenteeism, and overall work impairment. A logistic regression was used to analyze group differences on labor force participation.

All analyses applied sampling weights (calculated using data from the NHWS and the US Bureau of the Census) to project the results to the US population, though bivariate comparisons also include unweighted statistics (i.e., frequencies and means with applying a sampling weight). The *a priori* cutoff for statistical significance was set as $p < 0.05$. All analyses were conducted using SAS 9.1 (SAS Institute, Inc., Cary, NC, USA)

Results

Sociodemographic and health risk behavior differences

Table 1 compares respondent characteristics by the HCV status. Subjects with HCV diagnosis were older (50.4 vs. 46.0 years; $p < 0.0001$) and less likely to be female (41.9 vs. 52.2%; $p < 0.0001$) compared to those without HCV. Fewer HCV subjects were married (54.5 vs. 59.4%, $p = 0.017$) or college-educated (22.6 vs. 36.1%, $p < 0.0001$). The proportion of those with an annual income of less than \$25,000 was lower in the HCV group (31.8 vs. 18.9%, $p < 0.0001$) relative to controls. Those with HCV were more likely to smoke, less likely to have exercised in the past month, and less likely to currently consume alcohol. In addition, the prevalence of anxiety and depression was more than twice as high in the HCV group than in the control group.

Labor force participation

Using a regression modeling approach, HCV patients were found to be significantly less likely to be in the labor force than unmatched controls (unstandardized

Table 2. Predictors of labor force participation.

Predictor	<i>b</i>	OR	95% LCL	95% UCL	χ^2	<i>p</i> -value
Intercept	-1.07	—	—	—	295.06	<0.0001
Age 18–29	2.55	12.75	11.74	13.84	3681.70	<0.0001
Age 30–39	2.95	19.03	17.51	20.67	4843.14	<0.0001
Age 40–49	3.04	20.84	19.23	22.59	5484.81	<0.0001
Age 50–59	2.60	13.49	12.47	14.59	4242.89	<0.0001
Age 60–69	1.01	2.75	2.55	2.96	709.27	<0.0001
Age ≥70 (reference)	—	—	—	—	—	—
Female	-0.42	0.66	0.63	0.68	504.92	<0.0001
White (reference)	—	—	—	—	—	—
Hispanic	0.08	1.08	1.01	1.16	5.64	0.0176
Black	0.14	1.15	1.08	1.22	19.42	<0.0001
Asian	-0.15	0.86	0.79	0.94	11.14	0.0008
Other ethnicity	-0.03	0.97	0.87	1.07	0.41	0.5216
Married/living with partner	-0.42	0.65	0.63	0.68	439.37	<0.0001
College educated	0.57	1.77	1.70	1.84	762.31	<0.0001
Household income: <\$25 k	-1.08	0.34	0.32	0.36	1157.98	<0.0001
Household income: \$25 k to <\$50 k	-0.43	0.65	0.62	0.69	258.25	<0.0001
Household income: \$50 k to <\$75 k	-0.20	0.82	0.78	0.87	51.89	<0.0001
Household income: ≥\$75 k	—	—	—	—	—	—
Household income: decline to answer	-0.77	0.47	0.43	0.50	341.35	<0.0001
Health insurance	0.42	1.53	1.45	1.60	276.43	<0.0001
Currently smoke	0.07	1.08	1.03	1.12	10.63	0.0011
Currently exercise	0.09	1.10	1.06	1.14	24.53	<0.0001
Consume alcohol	0.41	1.50	1.45	1.56	468.10	<0.0001
BMI: underweight	-0.27	0.76	0.67	0.87	17.37	<0.0001
BMI: normal weight (reference)	—	—	—	—	—	—
BMI: overweight	0.12	1.13	1.07	1.18	25.04	<0.0001
BMI: obese	0.04	1.04	0.99	1.09	2.26	0.133
BMI: DECLINE to answer	0.01	1.01	0.89	1.15	0.02	0.8771
Comorbidity count	-0.07	0.93	0.93	0.94	762.37	<0.0001
Anxiety	0.01	1.01	0.96	1.08	0.22	0.6358
Depression	-0.10	0.91	0.86	0.96	11.19	0.0008
Hepatitis C	-0.32	0.73	0.62	0.85	15.29	<0.0001

b = unstandardized regression estimate; OR = odds ratio; 95% LCL = lower 95% confidence limit of the odds ratio; 95% UCL, upper 95% confidence limit of the odds ratio; χ^2 = chi-square test.

regression estimate (*b*) = -0.32, odds ratio (OR) = 0.73, *p* < 0.0001), even after adjusting for sociodemographic, health risk, and comorbidity variables, many of which were significantly associated with labor force participation (see Table 2). In particular, young age, male gender, Hispanic ethnicity, black ethnicity, unmarried status, college education, high household income, health insurance, smoking status, exercise, alcohol consumption, overweight BMI, fewer comorbidities, and lack of depression were all significantly predictive of labor force participation.

Work productivity loss

Among those who were currently employed (full-time, part-time, or self-employed), the effect of HCV was examined on absenteeism, presenteeism, and overall work impairment after adjusting for sociodemographics, health risk behaviors, and comorbidity status (see Table 3). Interestingly, those over the age of 70 generally had lower levels of impairment relative to other age groups. Although seemingly counterintuitive, an apparent self-selection bias may be present. Past retirement age, employment is less likely to be a necessity, thus workers over 70

still remain in the work force largely by choice and may be relatively healthier after adjusting for all other sociodemographic, health risk behavior, and comorbidity variables. White ethnicity, college education, and high household income were associated with lower levels of impairment. In regards to health risk behaviors and comorbidities, smoking, obesity, depression (though not anxiety), and a high comorbidity status were associated with more impairment while exercise was associated with less impairment.

Of primary interest, the effect of HCV was significantly associated with presenteeism (*b* = 0.21, *p* = 0.02), overall work impairment (*b* = 0.23, *p* = 0.01), and marginally related to absenteeism (*b* = 0.48, *p* = 0.05). After controlling for the set of covariates outlined above, patients with HCV reported absenteeism levels of 4.88% (compared with 3.03% for unmatched controls), presenteeism levels of 16.69% (compared with 13.50%), and overall work impairment levels of 19.40% (compared with 15.35%).

Activity impairment

Among all respondents, the effect of HCV was examined on activity impairment after adjusting for

Table 3. Generalized linear model regression estimates when predicting each of the work productivity and activity impairment metrics.

	Absenteeism	Presenteeism	Overall work impairment	Activity impairment
Intercept	0.04	1.16*	1.42*	2.49*
Age 18–29	0.84*	1.09*	1.00*	0.09*
Age 30–39	0.70*	0.97*	0.89*	0.04*
Age 40–49	0.45*	0.80*	0.71*	–0.02
Age 50–59	0.32	0.63*	0.55*	–0.05*
Age 60–69	0.00	0.38*	0.30*	–0.10*
Age ≥70 (reference)	–	–	–	–
Female	–0.07	–0.08*	–0.07*	–0.02
White (reference)	–	–	–	–
Hispanic	0.55*	0.24*	0.26*	0.14*
Black	0.39*	0.10*	0.14*	0.03
Asian	0.27*	0.40*	0.38*	0.27*
Other ethnicity	0.21	0.13*	0.14*	0.04
Married/living with partner	–0.02	0.00	0.00	0.03*
College educated	–0.16*	–0.01	–0.04*	–0.06*
Household income: <\$25 k	0.20*	0.20*	0.19*	0.31*
Household income: \$25 k to <\$50 k	0.13*	0.12*	0.12*	0.19*
Household income: \$50 k to <\$75 k	–0.04	0.07*	0.05*	0.10*
Household income: ≥\$75 k	–	–	–	–
Household income: decline to answer	0.16	–0.05	–0.01	0.10*
Health insurance	0.02	0.1*	0.07*	0.01
Current smoker	0.22*	0.15*	0.14*	0.12*
Regular exercise	–0.12*	–0.06*	–0.08*	–0.18*
Current drinker	–0.06	0.00	–0.01	–0.07*
BMI: underweight	0.48*	0.17*	0.20*	0.11*
BMI: normal weight (reference)	–	–	–	–
BMI: overweight	–0.02	–0.01	0.00	0.02
BMI: obese	0.11	0.11*	0.11*	0.18*
BMI: decline to answer	–0.03	–0.04	–0.05	0.17*
Comorbidity count	0.15*	0.13*	0.13*	0.11*
Anxiety	0.12	0.02	0.04	0.00
Depression	0.17*	0.25*	0.22*	0.24*
Hepatitis C	0.48	0.21*	0.23*	0.14*

* $p < 0.05$.

sociodemographics, health risk behaviors, and comorbidities (see Table 3). The same factors as noted above (ethnicity, education, income, etc.) were generally associated with impairment. Much like the other forms of work impairment, there was a significant effect of HCV on activity impairment ($b = 0.14$, $p < 0.01$). Respondents with HCV reported significantly higher levels of activity impairment (25.01%) relative to non-HCV subjects (21.78%), after adjusting for the set of covariates.

Propensity scoring

The research questions above were then examined using a propensity matching methodology to replicate the observed regression modeling effects. As expected, post-propensity matching there were no significant differences between the HCV group and the matched control group on any of the variables. Next, employed patients with HCV ($n = 293$) were compared to matched employed controls ($n = 293$). These groups were also generally comparable post-match. Although patients with HCV reported a

significantly higher rate of smoking (54.1 vs. 43.9%, $p = 0.04$), no other significant differences were observed.

Using a propensity match approach to estimate the effect of HCV on labor force participation revealed no difference between the groups. Both respondents with HCV and matched controls were equally likely to be in the labor force (52.85 vs. 54.54%, $p = 0.55$).

Even after accounting for the imbalance of smoking behavior, employed HCV patients reported significantly higher levels of absenteeism ($b = 0.64$, $p = 0.006$), presenteeism ($b = 0.26$, $p = 0.009$), and overall work impairment ($b = 0.30$, $p = 0.002$) relative to matched employed controls. Specifically, HCV patients reported absenteeism levels of 9.27% (compared with 4.87% for matched employed controls), presenteeism levels of 27.23% (compared with 21.00%), and overall work impairment levels of 31.57% (compared with 23.47%). When comparing all patients with HCV and matched controls, there was a significant effect of HCV status on activity impairment ($b = 0.17$, $p < 0.0001$). Levels of activity impairment were 47.91% among patients with HCV, compared with 40.22% among matched controls.

Discussion

The aim of this project was to document, using a nationally-representative sample, the effect of HCV on labor force participation and productivity loss. The regression results suggest that patients with HCV are significantly less likely to participate in the labor force and have significantly greater work productivity loss and impairment in daily activities. Using a propensity score methodology to replicate those findings, many of the regression results were corroborated. The lone exception was that labor force participation was not significantly lower in the HCV group using propensity score matching. In part, this may be due to the statistical power discrepancy. Because the regression model approach had a much larger control group, the power to detect small effects was enhanced. It is possible the effect of HCV on labor force is present but relatively modest (indeed, the trend was such that those with HCV had lower labor force participation rates). In contrast, the impact of HCV on productivity loss and activity impairment was detected in both methodologies, highlighting the magnitude of these effects.

These findings have important implications given the limited number of studies investigating the impact of HCV on labor force participation, absenteeism and presenteeism. Indeed, only one national study could be identified that assessed the impact of HCV on employment status⁹. This study found that positive HCV status with normal alanine aminotransferase (ALT) levels in males was associated with a 10.7% reduction in labor force participation while positive HCV status and elevated ALT was associated with a 17.5% reduction.

Our results also suggest a detrimental effect of HCV on labor force participation, though only when analyzed using a logistic regression approach. It is important to note that the Jacobs *et al.*⁹ analysis controlled for self-perceived health status but not a complete list of comorbidities as done in the present analysis. By demonstrating a significant effect even after controlling for comorbidities, anxiety, and depression, the incremental effect of HCV was more properly isolated (though, as noted above, this effect is likely small).

The literature on the impact of HCV on absenteeism and presenteeism is just as limited with only one study identified¹¹. Su *et al.* utilized employee records from multiple large employers in the US and found (via regression modeling) that employees with HCV had 1.85 times more absence days than controls¹¹. The number of annual absence days varied by type. The difference was smaller for those taking sick leave and for those on workers' compensation and greater for those on short-term or long-term disability. Overall, those with HCV recorded 4.15 more days of absence per employee than the control cohort. Productivity was lower with employees with HCV processing 7.5% fewer units per hour than controls. All healthcare

benefit costs among employees with HCV were significantly higher than the same costs among employees without HCV.

Our absenteeism results were consistent with Su *et al.*¹¹, in that levels of absenteeism were 1.61 times higher (4.88 vs. 3.03%) in our regression modeling approach. However, our presenteeism results were much stronger than Su *et al.*¹¹ in that impairment was 1.24 times higher (16.69 vs. 13.50%) in our regression modeling approach. The difference in effects may be due to the use of employer data versus our use of patient reports; the latter of which may be better suited to picking up more subtle effects of productivity, especially in cases when employee productivity is difficult for employers to quantify. It is also important to note that the Su *et al.*¹¹ paper used linked employer and claims data, which would have neglected patients whose physician did not code their HCV diagnosis. Further, the sample was skewed towards full-time employees. The Su *et al.*¹¹ study also neglected those not currently employed. Indeed, the current study was the first of its kind to demonstrate the impact of HCV infection on impairment outside of the workplace.

Limitations

All HCV diagnoses and work productivity measures were patient-reported and may have introduced measurement error. Although the results from the propensity scoring and regression modeling generally coincided in demonstrating the burden of HCV on work productivity, it is possible that there may be additional variables not included, which could explain the observed differences in health outcomes. This is an important limitation. However, it should be noted that the most likely alternative explanations for the relationship between HCV status and work productivity (such as comorbidities, health behaviors, etc) have been accounted for in both a regression modeling and propensity score matching framework. The Internet survey methodology represents another limitation, as certain disenfranchised groups without Internet access would not have been able to participate. Naturally, this study only includes respondents from the US and it is not clear whether these effects generalize to other geographies. It is also important to note that the study was cross-sectional and likely included a mix of HCV patients, from those newly diagnosed to those who are in the later stages of infection. It would be informative, particularly for some employers with a short-term focus, to use a time series approach to document work productivity changes pre- and post-HCV diagnosis. Although using both a regression approach and a propensity score matching approach provided a robust series of results, other methodologies (such as a multi-part model) may be useful, particularly when estimating specific costs associated with HCV.

Conclusions

Although much of the work in the HCV area has focused on direct costs, our results suggest the indirect costs should not be ignored when quantifying the societal burden of HCV. To our knowledge, this is the first study which has utilized a large, nationally-representative data source for identifying the impact of HCV on labor force participation and work and activity impairment using both a propensity-score matching and regression modeling framework. Results from both methodologies converged to indicate that HCV is associated with persistent indirect economic costs from a societal perspective.

Transparency

Declaration of funding

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Both M.D. and J.-S.W. of Kantar Health while Y.Y. and G.L. are employees of Bristol-Myers Squibb. P.L. served as a consultant to Kantar Health.

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