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Seroprevalence of markers of hepatitis C virus exposure and associated factors in adults aged 18-39 years in the Arctic Russian city of Arkhangelsk: a cross-sectional study

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ABSTRACT

Hepatitis C, caused by the hepatitis C virus (HCV), is a major public health issue in Russia. The aim of our study was to assess the seroprevalence of markers of HCV exposure and factors associated with HCV seropositivity among the general population aged 18–39 years in the city of Arkhangelsk, Northwest Russia. A social research agency applied a quota sampling method to recruit study participants using cell phone numbers. All participants (n = 1243) completed a self-administered questionnaire and provided a blood sample. Sixty-five participants (5.2%, 95% confidence interval [CI] 4.9–5.5) tested positive for HCV IgM+G antibodies, and of these, 55 (84.6%) did not know that they were exposed to HCV. In multivariable logistic regression analysis, HCV seropositivity was significantly associated with older age, a history of injecting drug use, and having ever received a blood transfusion. To reach the goal of the World Health Organisation's Global Health Sector Strategy on Viral Hepatitis, regional preventive programmes should include measures to reduce injecting drug use as well as scaling up harm-reduction and treatment programs for drug addicts.

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KEYWORDS

Hepatitis C; HCV; seroprevalence; associated factors; injecting drug use; Russia; population-based study

Introduction

The prevalence of hepatitis C virus (HCV) infection is high worldwide [1]. HCV is a blood-borne virus which can be transmitted through injecting drug use (IDU), transfusion of infected blood, and reuse or inadequate sterilisation of medical equipment. It can also be transmitted from mother to child and in rare cases during unprotected sex [2]. HCV can cause acute hepatitis, but about 80% of the infected people do not develop symptoms [3]. From 60% to 80% of HCV-infected persons develop chronic hepatitis C, which can be asymptomatic for decades, but may lead to cirrhosis or hepatocellular carcinoma [4]. Globally, more than 71 million people have chronic HCV infection, and about 400,000 people die annually due to HCV infection [3]. Worldwide, the countries with the highest number of HCV cases are China, Pakistan, India, Egypt and Russia [5]. At present, there is no vaccine against HCV [3]. Although new antiviral medicines can cure more than 95% of HCV- infected people, access to diagnostic procedures and treatment is still low on the global scale [6].

In 2016, the World Health Assembly adopted the first Global Health Sector Strategy on Viral Hepatitis, with the global aim of reducing the incidence of HCV infection by 90% and mortality due to HCV infection by 65% by 2030. The World Health Organisation (WHO) proposed 10 core indicators to support the implementation and monitoring of the strategy [7]. The WHO Regional office for Europe assessed the availability of the data for each core indicator in the region and came to the conclusion that current data sources in most European countries are insufficient and that seroprevalence surveys are required to determine the size of the undiagnosed population [8].

HCV infection is an important public health issue in Russia [9]. In 2018, the incidence of reported acute hepatitis C was 1.1 per 100,000 while the incidence of reported chronic hepatitis C was 32.7 per 100,000 (Figure 1) [10]. About 60% of chronic hepatitis C cases

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Figure 1. Incidence of hepatitis C (HC) in Russia and the Arkhangelsk region in 1996–2018 (per 100,000 population).

are diagnosed among adults aged 20–39 years [11]. Russia has a national strategy for preventing HCV infection. All Russian citizens have the right to treatment for HCV infection free of charge. Moreover, HCV testing is free of charge for blood donors, pregnant women, and recipients of blood transfusions. HCV testing is obligatory for health-care workers and patients who are considered to be at increased risk for viral hepatitis [12]. People who inject drugs are eligible for free of charge treatment in public health-care facilities. The total economic burden of hepatitis C in Russia was estimated to be 48 billion Russian rubles (~0.83 billion US dollars) in 2010 and continues to increase [13].

The only study on the seroprevalence of HCV in Russia conducted the last 20 years was published in Russian, and is unavailable to the international audience [14]. This study was conducted in 2008 in five Russian regions (Moscow, Rostov, and Sverdlovsk Regions, as well as the Tyva and Sakha (Yakutia) Republics). All these settings are situated thousands of kilometres away from the Arkhangelsk Region where our study was performed. The overall seroprevalence of HCV among the participants of all ages in the study was 2.6% [14].

The aim of our study was to assess the seroprevalence of markers of HCV exposure and factors associated with HCV seropositivity among adults aged 18–39 years in the city of Arkhangelsk, Northwest Russia. Further, to determine the proportion of that population with undiagnosed HCV infection, thus providing key information to health authorities needed to scale up interventions proposed by the WHO's Global Health Sector Strategy on Viral Hepatitis in this part of Russia [8].

Materials and methods

Study population and study design

This study was a part of a large Russian-Norwegian project [15], and it drew its study sample from the general population aged 18–39 in the Arctic Russian city of Arkhangelsk using a guota sampling method [16]. Arkhangelsk has a population of approximately 350,000 people, and the official data show that its population has an incidence of acute (1.48 per 100,000 in 2018) and chronic (40.8 per 100,000 in 2018) hepatitis C that is close to the national average. A social research agency recruited participants using cell phone numbers. All participants completed a self-administered questionnaire and provided a blood sample between September 2010 and June 2011. Detailed information on enrolment procedures, questionnaire data, and sample collection has been described elsewhere [15]. Altogether, 1265 adults participated in the study. Twenty-two of them had missing blood test results and were excluded from the analyses. Thus, the final study sample consisted of 1243 participants.

Description of the variables

The participants provided information on demographic factors, socioeconomic factors, smoking status, alcohol use, sexual and preventive behaviour, history of sexually transmitted infections (STIs)/HIV, injecting drug use (IDU), blood donation, blood transfusions, operations, tattoos, and whether they knew anyone with hepatitis C. Participants reported their family income on a scale of 1 to 10, based on how they thought their income compared

to the average income in Arkhangelsk. We then categorised these responses as low (1–4), medium (5–7), and high (8–10) income. Having tattoos was dichotomised as "yes" and "no". The variable "knowing someone with hepatitis C" was divided into "no", "yes", and "do not know". The description of other variables used in this study can be found elsewhere [17].

Serological testing and reporting of test results

Serum samples were sent daily to the research laboratory of the Northern State Medical University (NSMU), where they were centrifuged and frozen at -20°C. Once a month, all untested samples were transported to the laboratory of the Regional STI clinic in Arkhangelsk, where serological tests for HCV IgM+G, were done by enzyme-linked immunosorbent assay, according to the manufacturer's instructions. A confirmation test was performed in case of positive results on IgM+G. The sample was tested for antibodies to HCV antigens corresponding to the parts of proteins encoded by the structural (core) and non-structural (NS3, NS4, NS5) genome parts. If a participant had positive results for one or both tests (IgM+G or IgM), he/she was considered to have been exposed to HCV and was categorised as seropositive. All reagents were produced by the Vector-Best company, Novosibirsk, Russia.

All participants were asked to provide a contact e-mail address or telephone number so they could be informed on their test results and the results were reported via e-mail or cell phones. The participants with positive test results were referred to a specialist in infectious diseases who reported back that 17 of them complied. Further follow-up and RNA testing of positive participants was not part of this study as these participants were referred to the health service.

Statistical analyses

The direct standardisation technique was applied to provide the age-standardised seroprevalence of markers of HCV infection to the 2010 Arkhangelsk population, based on the 2010 census. Associations between HCV seropositivity and independent variables were studied using crude logistic regression. All associations with p < 0.15 or lower in crude analyses were included in the multivariable model [18]. Crude and adjusted odds ratios with 95% confidence intervals (CIs) were estimated. All analyses were performed using Statistical Package for Social Sciences version 20 (SPSS, Inc., Chicago, IL, USA).

Ethical approval

This study was conducted in accordance with the Helsinki Declaration and approved by the Ethical Committee at the NSMU in the city of Arkhangelsk (Protocol 04/03 from 15 March 2010). The participants were given an invitation letter with information about the study aims, procedures, and a telephone number they could call to obtain their test results. All participants signed a written informed consent form.

Results

The final study sample consisted of 543 men and 700 women (mean age 27.6, standard deviation [SD] = 5.6 years and 27.0, SD = 5.7 years, respectively). Most of the participants had high education (58.9%) and medium income (72.0%). Ever smokers comprised 60.5% of the study sample, and 80.5% of participants reported at least one episode of binge drinking (at least 6 units of alcohol in one occasion) (Table 1). Previous HIV testing was reported by 55.4% of men and 73.9% of women. No one reported being diagnosed with HIV infection (data not shown). A history of IDU was reported by 2.7% (n = 34) of the sample, and of those, 14 were seropositive (10 men and 4 women). More than one-fifth of the sample had ever donated blood; 4.7% had ever received a blood transfusion. Among the participants, 12.1% knew someone with hepatitis C, and 16.9% did not know if they knew someone with hepatitis C (Table 1).

Laboratory results revealed that 65 participants (5.2%, 95% CI 4.9–5.5) tested positive for HCV IgM+G antibodies, among whom six tested positive for IgM, only. The seroprevalence of markers of HCV exposure was 6.4% (95% CI 4.5–8.9) among men and 4.3% (95% CI 2.9–6.1) among women. Fifty-five of the 65 seropositive participants (84.6%) did not know that they were exposed to HCV; eight (12.3% of seropositive participants) had been previously told by medical workers that they had HCV infection, and two persons (3.1%) did not know whether or not they had ever had HCV infection.

Compared to the group-level census data from Arkhangelsk in the same period, women and younger individuals were slightly overrepresented in our study sample. After direct standardisation by age, the overall seroprevalence of markers of HCV exposure increased to 5.6% (95% CI 5.3–5.9). It increased to 7.1% (95% CI 6.7–7.5) among men, but decreased to 4.2% (95% CI 3.8–4.6) among women (Table 1). So, the sex difference in HCV seropositivity became significant after direct age standardisation.

In crude logistic regression, HCV seropositivity was significantly associated with older age, age at sexual

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Table 1. Crude	odds ratios (OR)) and 95% confiden	ce intervals (CI) for	the association	between seropre	valence of markers of
hepatitis C virus	s (HCV) exposure	and its correlates an	nong 1243 participa	nts aged 18–39	years, 2010–2011,	Arkhangelsk, Russia.

Variable	N (%)	N seropositive for HCV (%)	Crude OR (95% CI)
Sex			
Men	543 (43.7%)	35 (6.4%)	referent
Women	700 (56.3%)	30 (4.3%)	0.65 (0.39-1.07)
Age			
18–24 years	497 (40%)	18 (3.6%)	referent
25–29 years	296 (23.8%)	12 (4.1%)	1.12 (0.53–2.37)
30–34 years	278 (22.4%)	20 (7.2%)	2.06 (1.07–3.97)
35–39 years	172 (13.8%)	15 (8.7%)	2.54 (1.25–5.16)
Marital status			
Single	571 (45.9%)	28 (4.9%)	referent
Divorced/widowed	86 (6.9%)	7 (8.1%)	1./2 (0./3-4.0/)
Conabiting	213 (17.1%)	II (5.2%)	1.06 (0.52-2.16)
Married	373 (30.0%)	19 (5.1%)	1.04 (0.57-1.89)
	200 (16 90/)	12 (6 20%)	referent
Average (secondary/vocational)	209 (10.0%)	15 (0.270)	0.84 (0.40 - 1.70)
High (incomplete higher/higher)	302 (24.3%) 732 (58.0%)	10 (J.570) 36 (4 0%)	0.64 (0.40 - 1.79) 0.78 (0.41 - 1.50)
Income	752 (50.570)	50 (1.270)	0.70 (0.1-1.50)
Low	207 (16 7%)	7 (3.4%)	referent
Medium	895 (72.0%)	52 (5.8%)	1.76 (0.79–3.94)
High	141 (11.3%)	6 (4.3%)	1.27 (0.42–3.86)
Smoking status			(
Never	491 (39.5%)	21 (4.3%)	referent
Ever	752 (60.5%)	44 (5.9%)	1.39 (0.82–2.37)
Binge drinking			
Never	242 (19.5%)	12 (5.0%)	referent
Ever	1001 (80.5%)	53 (5.3%)	1.07 (0.56–2.04)
Age at sexual debut			
≥18 years	517 (41.6%)	16 (3.1%)	referent
<18 years	726 (58.4%)	49 (6.7%)	2.27 (1.27–4.03)
N of sexual partners in the last 6 months	044 (75 70()	12 (1 50()	<i>c</i> .
U and I	941 (75.7%)	42 (4.5%)	referent
≥∠ Lifetime N of council nontrone	302 (24.3%)	23 (7.6%)	1.// (1.04–2.99)
0 and 1	124 (10 904)	5 (2 704)	roforont
	134 (10.0%)	5 (5.7%) 11 (2.6%)	
2-J 6 and more	685 (55.1%)	49 (7.2%)	1 99 (0.23-2.01)
Condom use during last sexual encounter	005 (55.170)	49 (7.270)	1.55 (0.76-5.65)
Yes	339 (27,3%)	23 (6.8%)	referent
No and don't remember	904 (72.7%)	42 (4.6%)	0.67 (0.40–1.13)
Condom use with causal partner			(
Always, without causal partners	1112 (89.5%)	57 (5.1%)	referent
Sometimes, rare, never	131 (10.5%)	8 (6.1%)	1.20 (0.56-2.58)
Sexual relations with a partner of the same sex			
Never	1197 (96.3%)	61 (5.1%)	referent
Ever	46 (3.7%)	4 (8.7%)	1.77 (0.62–5.11)
Sexual relations with a person reporting injecting drug use			
No	1094 (88.0%)	47 (4.3%)	referent
Yes	45 (3.6%)	8 (17.8%)	4.82 (2.13-10.92)
Don't know	104 (8.4%)	10 (9.6%)	2.37 (1.16–4.84)
Nover ^{§§}	006 (90 104)	40 (4 00%)	roforont
Vec	990 (00.1%) 247 (10.0%)	49 (4.9%) 16 (6.5%)	1 34 (0 75_2 40)
Injecting drug use	247 (19.970)	10 (0.570)	1.54 (0.75-2.40)
Never	1209 (97.3%)	51 (4.2%)	referent
Ever§	34 (2.7%)	14 (41.2%)	15.89 (7.60-33.26)
Donated blood			
Never	950 (76.4%)	53 (5.6%)	referent
Ever	293 (23.6%)	12 (4.1%)	0.72 (0.38-1.37)
Received a blood transfusion			
Never	1184 (95.3%)	57 (4.8%)	referent
Ever	59 (4.7%)	8 (13.6%)	3.10 (1.41–6.84)
Operations			
Never	739 (59.5%)	34 (4.6%)	referent
Ever Total a	504 (40.5%)	31 (6.2%)	1.36 (0.82–2.24)
	1071 (06 20/)	40 (4 (0))	
NU Voc	10/1 (80.2%)	49 (4.6%) 16 (0.2%)	
Tes Knowing someone with honotitis C	172 (13.8%)	10 (9.3%)	2.14 (1.19-3.86)
No	882 (71 00%)	36 (4 10%)	rafarant
Yes	150 (12 1%)	16 (10 7%)	2.81 (1 52-5 20)
Don't know	210 (16.9%)	13 (6.2%)	1.55 (0.81–2.98)

STI: sexually transmitted infections; HSV-2: herpes simplex virus type 2 $^{\$I}$ Included 11 who didn't want to answer $^{\$\$}$ Included 9 who didn't want to answer

Table 2. Hepatitis C virus (HCV) seropositivity and correlates by multivariable logistic regression analysis among 1243 persons in a population-based seroepidemiological study of adults aged 18–39 years, 2010–2011, Arkhangelsk (Russia).

VariableAdjusted OR95% ClAgeReferent $18-24$ yearsReferent $25-29$ years 0.87 $0.38-2.03$ $30-34$ years 1.80 $0.88-3.68$ $35-39$ years 2.55 $1.18-5.50$ Age at sexual debut 218 yearsReferent $≤17$ years 1.72 $0.91-3.25$ N of sexual partners in the last 6 months 0 and 1Referent $≥2$ 1.66 $0.93-2.97$ Sexual relations with a person reporting injecting drug use NoReferentNoReferentYes 2.22 $0.80-6.13$ Don't know 1.51 $0.69-3.34$, , , , , , , , , , , , , , , , , , , ,		
Age Referent 18–24 years Referent 25–29 years 0.87 0.38–2.03 30–34 years 1.80 0.88–3.68 35–39 years 2.55 1.18–5.50 Age at sexual debut 2.55 1.18–5.50 Age at sexual debut 2.55 1.18–5.50 Age at sexual debut 2.55 1.72 ≥18 years Referent 517 years 0 and 1 Referent 22 1.66 0.93–2.97 Sexual relations with a person reporting injecting drug use Ves No Referent Yes 2.22 0.80–6.13 Don't know 1.51 0.69–3.34	Variable	Adjusted OR	95% Cl
18–24 years Referent 25–29 years 0.87 0.38–2.03 30–34 years 1.80 0.88–3.68 35–39 years 2.55 1.18–5.50 Age at sexual debut 2.55 1.18–5.50 All sears Referent 51 ≤17 years 1.72 0.91–3.25 N of sexual partners in the last 6 months 0.93–2.97 Sexual relations with a person reporting injecting drug use 0.93–2.97 No Referent Ves 2.22 0.80–6.13 Don't know 1.51 0.69–3.34	Age		
25-29 years 0.87 $0.38-2.03$ $30-34$ years 1.80 $0.88-3.68$ $35-39$ years 2.55 $1.18-5.50$ Age at sexual debut \geq 15 years 867 ≥ 18 years Referent \leq 17 years ≤ 17 years 1.72 $0.91-3.25$ N of sexual partners in the last 6 months 0 and 1 Referent ≥ 2 1.66 $0.93-2.97$ Sexual relations with a person reporting injecting drug use No Referent No Referent Yes 2.22 $0.80-6.13$ Don't know 1.51 $0.69-3.34$	18–24 years	Referent	
$30-34$ years 1.80 0.88-3.68 $35-39$ years 2.55 1.18-5.50 Age at sexual debut 2.55 1.18-5.50 ≥ 18 years Referent 2.55 ≤ 17 years 1.72 0.91-3.25 N of sexual partners in the last 6 months 0 and 1 Referent ≥ 2 1.66 0.93-2.97 Sexual relations with a person reporting injecting drug use No Referent No Referent 2.22 0.80-6.13 Don't know 1.51 0.69-3.34	25–29 years	0.87	0.38-2.03
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Age at sexual debut ≥ 18 yearsReferent ≤ 17 years 1.72 $0.91-3.25$ N of sexual partners in the last 6 monthsReferent0 and 1Referent ≥ 2 1.66 $0.93-2.97$ Sexual relations with a person reporting injecting drug use NoReferentNoReferentYes 2.22 $0.80-6.13$ Don't know 1.51 $0.69-3.34$	35–39 years	2.55	1.18-5.50
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months Referent 0 and 1 Referent ≥2 1.66 0.93–2.97 Sexual relations with a person reporting injecting drug use V No Referent V Yes 2.22 0.80–6.13 Don't know 1.51 0.69–3.34	N of sexual partners in the last 6		
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Yes2.220.80-6.13Don't know1.510.69-3.34	No	Referent	
Don't know 1.51 0.69–3.34	Yes	2.22	0.80-6.13
	Don't know	1.51	0.69-3.34
Injecting drug use	Injecting drug use		
Never Referent	Never	Referent	
Ever 8.52 3.66–19.83	Ever	8.52	3.66-19.83
Received a blood transfusion	Received a blood transfusion		
Never Referent	Never	Referent	
Ever 3.19 1.33–7.66	Ever	3.19	1.33–7.66
Tattoo	Tattoo		
No 1.29 0.65–2.56	No	1.29	0.65–2.56
Yes	Yes		
Knowing someone with HCV	Knowing someone with HCV		
infection	infection		
No Referent	No	Referent	
Yes 1.66 0.82–3.36	Yes	1.66	0.82-3.36
Don't know 1.50 0.75–3.02	Don't know	1.50	0.75-3.02

OR: odds ratio, CI: confidence interval

debut <18 years, having two or more sexual partners in the last 6 months, having ever received a blood transfusion, having a tattoo, and knowing someone with hepatitis C (Table 1). The highest seroprevalence of markers of HCV infection and the strongest association with HCV seropositivity were found among participants with a history of IDU and those with sexual partners who had a history of IDU. In total, 45 persons reported having had a sexual partner with a history of IDU, 10 (22%) of whom had a history of IDU themselves. One hundred four participants reported that they did not know the HCV serostatus of their partners. Marital status, education, income, smoking status, binge drinking, self-reported STIs, and ever having donated blood were not associated with HCV seropositivity. Women had lower odds of being seropositive for HCV than men. In the multivariable logistic regression analysis, HCV seropositivity was significantly associated with older age, history of IDU and history of blood transfusions (Table 2).

Discussion

The seroprevalence of markers of HCV infection among adults aged 18–39 years in the general population in

this study was 5.2%. This is twice as high as the seroprevalence observed in Eastern Europe for the general adult population (3.2% in Romania, 2.4% in Latvia, and 2.0% in Slovakia). These countries did not experience economic collapse to the same degree as Russia with a dramatic increase in IDU and cuts in health-care expenditures. At the same time, a seroprevalence of HCV of 5.9% has been reported in Italy which is similar to our findings. Considerable variations in the prevalence of HCV across Europe can be explained by different transmission routes, risk factors and prevention strategies used in different countries [19].

Our data are in a disagreement with data on the seroprevalence of markers of HCV infection among pregnant women in the Arkhangelsk region (0.6% in 2012) [20]. We propose two explanations for this. First, according to the official statistics the incidence of HCV infection is higher in the city of Arkhangelsk than in the rural areas and small towns in the region [10]. Second, data from Europe suggest that pregnant women are not a reliable population to use when trying to estimate the seroprevalence HCV in the general population. Indeed, a systematic review from 2017 reported that the seroprevalence of HCV among pregnant women was several times lower than that in the general population in European countries [19]. Data from our study are close to data from another Russian study published in 2001, which reported a seroprevalence of markers HCV infection of 5.3% among adults aged 25-64 years in Siberia [21]. However, our 30-39-year-old participants had a higher seroprevalence of markers of HCV exposure compared to those aged 20-29 years in Siberia (7.8% versus 3.8%). Data from another Russian study showed a seroprevalence of HCV of 3.1% among 20-29 year-olds and 4.9% in 30-39 year-olds [14]. This pattern may be explained by the high incidence of acute hepatitis C among 15-29 year-olds in 1997-2002 [10]. Our participants were aged 30-39 years in 2010-2011, thus they were 15-29 old in 1997-2002, when the incidence of HCV infection was high due to the IDU epidemic in the country [9]. The highest prevalence of HCV infection have been reported from Central Asia (3.6%) and Eastern Europe (3.3%) followed by Central Sub-Saharan Africa (2.1%) and North Africa/Middle East (1.7%). The prevalence of HCV in Central- and Western Europe, North- and Latin America does not exceed 1% [5].

Globally, the prevalence of anti-HCV antibody is greater among men, but the findings vary between countries. A study among blood donors in China did not find gender differences in the prevalence of anti-HCV antibody [22]. However, the prevalence of HCV infection among men was about twice as high as among women in the USA and Iran [23–25]. In our study, the prevalence of anti-HCV antibody was higher among men compared to women (6.4% vs. 4.3%). This may partly be explained by more IDU among men than among women. Different susceptibilities to the infection, different biological responses on HCV infection in men and women as well as social and behavioural factors were proposed to explain gender differences [26]. The dominant rote of transmission in the community can also partly explain existing gender differences. Further investigation of factors explaining gender differences in the seroprevalence of HCV markers may contribute to the development of gender-specific preventive strategies.

In our study, a history of IDU showed the strongest association with HCV seropositivity. This is in line with the findings from the USA and Western Europe. People with a recent history of IDU comprise 8.5% of all HCV infections globally with the greatest proportions in North America (30.5%), Latin America (22.0%) and Eastern Europe (17.9%). The prevalence of HCV infection among people with a recent history of IDU is 39.2% globally being the highest in Eastern Europe (48.6%), the Caribbean (47.6%) and Latin America (46.4%). Proportion of people with a recent history of IDU among the total population with HCV infection is the highest in North America (30.5%), Latin America (22.0%), Western- (17.9%) and Eastern (17.2%) Europe [27]. In our study, people who reported a history of ever IDU had a HCV seroprevalence of 41.2%. Several studies have shown a seroprevalence of markers of HCV infection up to 80% among individuals with a history of IDU in Russia [28-31]. According to the official data, IDU was responsible for 40% of acute hepatitis C cases in 1997 and for 21.5% of all new cases of acute hepatitis C in 2010 in Russia [32]. In the Arkhangelsk Region, 41.7% (5 out of 12) of registered acute hepatitis C cases were attributed to IDU in 2010 [20].

Drug possession is a crime in Russia; therefore some HCV-infected participants could provide false negative answers on their history of IDU leading to an underestimation of this route of transmission and potentially biasing our associations toward the null. There is no national programme to promote safe IDU in Russia, and despite the effectiveness of opioid substitution therapy and syringe exchange programmes in some countries [33,34], such programmes are neither implemented in the Arkhangelsk region nor in Russia in general. The number of beds in rehabilitation facilities for people with substance abuse disorders has increased in the last decade; however, there is still a lack of specialists in the field of addiction as well as equipment and facilities for preventive and harm-reduction programmes in the region [35].

From 2001 all donated blood has been screened for HCV as a part of the national infection control policy for blood

banks [36]. In our analyses, participants with self-reported blood transfusions had a higher risk of HCV seropositivity. According to Order 193 07.05.2003, "About implementation of quarantine of fresh frozen plasma in practice of work of blood service in the Russian Federation", all blood from donors had to be guarantined for 6 months and could only be used after the donor was tested for a second time. Obviously, it took time to reorganise the blood donation programme. We did not ask our participants to give information on when they received blood transfusions. Unsafe medical procedures have been reported as the most common risk factor for HCV infection in India and China [37,38]. Having a blood transfusion before 1991 is the second most common risk factor for HCV infection in Europe [39]. In the USA, the risk of HCV infection after blood transfusion decreased from 33% in the 1960s to 0.3% in the mid-1990s after the universal screening of blood and donors was implemented [40].

The proportion of cases of acute Hepatitis C attributed to medical procedures decreased from 12.8% in 1997 to 2.8 in 2010 in Russia. The corresponding numbers for Northwest Russia were 24.4% and 2.1% ([32], Figure 1). Taking into account that approximately 25% of people naturally clear the infection and 75% remain chronically infected after the exposure to HCV [41] the estimates for the prevalence of chronic HCV infection among our participants would be 5.3% (95% CI 5.0-5.6) among men and 3.2% (95% CI 2.9–3.5) among women. Our results suggest that almost 85% of the participants with markers of exposure to HCV infection had not previously received a formal diagnosis from medical personnel. Taking into account the same routes of transmission, our participants were asked about HIV testing, and no one reported HIV infection. Nevertheless, the HIV epidemic in the Arkhangelsk region is scaling up; the incidence rate has risen from 10.6 to 27.7 per 100,000 during the last 5 years, and IDU is considered to be responsible for at least 40% of all new HIV cases in the region [42]. Many researchers support the idea that Russia urgently needs to implement harmreduction programmes to prevent blood-borne infections among people who practice IDU [43–45]. Data on needle/ syringe distribution is one of the core indicators of the WHO monitoring and evaluation framework for HCV elimination in 2016–2021, but this indicator is not available in most regions of Russia.

Sexual transmission of HCV is possible but occurs seldom while the sexual transmission is a common route of transmission of hepatitis B virus and HIV [46,47]. Our study shows a low risk of sexual transmission of HCV. According to our data, the risk of being HCV seropositive among participants with two or more sexual partners during 6 months prior to the study was the same as for those who knew someone with hepatitis C, although neither of these results were significant. To our opinion, a relatively high rate of HIV testing in the study sample (55.4% of men and 73.9% of women) does not mean a selection of risk groups for our study. According to Russian Sanitation Rules 3.1.5.2826–10 "Prevention of HIV infection" testing for HIV infection is free of charge for all citizens. Moreover all blood donors, pregnant women and their sexual partners are tested for HIV during antenatal screening while military personnel, hospitalized patients before operations, people with Hepatitis B and C and other groups during routine medical check-ups. About 22% of the Russian population is tested for HIV every year [48].

Strengths and weaknesses of the study

Due to the cross-sectional design, a temporal relationship between high-risk behaviour and HCV seropositivity could not be established. Our sampling method cannot rule out selection bias associated with recruitment via cell phone owners. Even though the number of cell phones in Arkhangelsk region was high in 2011 (1890/1000 population, according to information from the Territorial Body of the Federal State Statistics Service of the Arkhangelsk region), generalisation of our study results should be done with caution. Moreover, as we investigated the adult population aged 18-39 years, we cannot generalise our results to the whole adult population of Arkhangelsk. In addition, the study questionnaire contained sensitive questions, which may have caused social desirability bias due to underreporting of high-risk behaviour [49]. However, more people at risk for HCV infection may have chosen to attend our study to avoid the stigmatisation they may have faced at the ordinary clinics. Blood sampling, seen as unpleasant by some, may have been a deterrent to participation [50]. The main strength of our study is its population-based design, including both sexes and presentation of sexspecific prevalence estimates. Other advantages of this study are its sufficient sample size, the use of multivariable statistics and adjustment for the most common potential confounders. We chose to study adults aged 18-39 years because that group is at highest risk for HCV infection [32]. To our knowledge, this study provides the most updated information of HCV seroprevalence and associated factors among young adults in an Arctic Russian urban setting in the international peer-reviewed literature.

Conclusions

We observed a rather high seroprevalence of HCV among the general population aged 18–39 years in Arkhangelsk. Factors associated with HCV seropositivity were included increasing age, IDU, and having received a blood transfusion. HCV screening programmes in population groups considered to be at risk for HCV infection should be expanded, i.e. among persons practicing IDU, persons acquiring tattoos or being subject to medical procedures/ blood transfusions abroad in places with unsafe hygienic standard and testing routines. To reach the goal of the World Health Organisation's Global Health Sector Strategy on Viral Hepatitis in reducing the incidence of and mortality due to HCV infection in the next decade, regional preventive programmes should include measures to reduce IDU as well as harm-reduction and treatment programmes for drug addicts.

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Conflict of interest

The authors report no conflicts of interest.

Data availability statement

Due to Russian legal regulation, the study data cannot be made available for public use. However, anonymized original data can be provided on request to experts if there are questions about statistical analyses. Public availability of the data was not among the issues that were cleared with the ethical committee at Northern State Medical University in Arkhangelsk. The authors are not authorized to make the data freely available because there is no informed consent for this from the participants (Federal Law No. 149-FZ on Information, Information Technologies and Data Protection 2006 (Data Protection Act)).

Authors' contributions

EK, TB, and AMG created the study design, all authors contributed to the research methodology. TB carried out data collection and data entry. EK, AMG, OS, and AS supervised data collection and data entry. TB analysed the data, and EK, AMG, and OS 8 👄 T. BALAEVA ET AL.

supervised the interpretation of the data. TB drafted the first version of the manuscript, all authors critically reviewed, commented on, and revised the manuscript, as well as approved the final version of the manuscript for submission.

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