RESEARCH ARTICLE

Understanding HIV Risk Behavior among Tuberculosis Patients with Alcohol Use Disorders in Tomsk, Russian Federation

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Abstract

Russian Federation’s (RF) HIV epidemic is the fastest growing of any country. This study explores factors associated with high HIV risk behavior in tuberculosis (TB) patients with alcohol use disorders in Tomsk, RF. This analysis was nested within the Integrated Management of Physician-delivered Alcohol Care for TB Patients (IMPACT, trial number NCT00675961) randomized controlled study of integrating alcohol treatment into TB treatment in Tomsk. Demographics, HIV risk behavior (defined as participant report of high-risk intravenous drug use and/or multiple sexual partners with inconsistent condom use in the last six months), clinical data, alcohol use, depression and psychosocial factors were collected from 196 participants (161 male and 35 female) at baseline. Forty-six participants (23.5%) endorsed HIV risk behavior at baseline. Incarceration history (Odds Ratio (OR) 3.93, 95% confidence interval (CI) 1.95, 7.95), age under 41 (OR: 2.97, CI: 1.46, 6.04), drug addiction (OR: 3.60 CI: 1.10, 11.77), history of a sexually transmitted disease (STD) (OR 2.00 CI: 1.02, 3.90), low social capital (OR: 2.81 CI: 0.99, 8.03) and heavier alcohol use (OR: 2.56 CI: 1.02, 6.46) were significantly more likely to be associated with HIV risk behavior at baseline. In adjusted analysis, age under 41 (OR: 4.93, CI: 2.10, 11.58), incarceration history (OR: 3.56 CI: 1.55, 8.17) and STD history (OR: 3.48, CI: 1.5, 8.10) continued to be significantly associated with HIV risk behavior. Understanding HIV transmission dynamics in Russia remains an urgent priority to inform strategies to address the epidemic. Larger studies addressing sex differences in risks and barriers to protective behavior are needed.
Introduction

In many settings where the HIV epidemic is driven by sexual transmission, alcohol use is an important risk factor. Research in many populations has linked alcohol use with HIV transmission via several mediating mechanisms, including psychosocial and socio-economic factors, as well as behavioral and biological risk factors. Underlying socio-economic vulnerability and instability are driving forces behind the convergence of alcohol use disorders (AUDs) and HIV, disproportionately affecting marginalized populations, such as the unemployed and incarcerated; the same populations also at increased risk for tuberculosis (TB). Heavy alcohol use affects high-risk sexual behavior in a number of ways, including its association with lower levels of condom use and/or multiple sex partners. This association has been demonstrated in diverse populations such as gay men [1] and men who have sex with men in Moscow, Russia, [2] adolescents, [3] men and women in East and Southern Africa, [4] and people with HIV in the Russian Federation. [5] Heavy alcohol use is also associated with immunomodulation and impaired host defense, which may increase the risk of infection and disease progression among individuals exposed to HIV or tuberculosis. Heavy alcohol use has also been associated with decreased adherence to TB and HIV treatment, [6] reduced viral load suppression [6] and death in HIV-infected patients [7] (particularly in women) and TB patients. [8] Conversely, HIV diagnosis may also influence alcohol use: in a Russian study from 1996, diagnosis with HIV was associated with a later diagnosis of an alcohol use disorder in 22% of the study’s participants. [9]

The epidemic in the Russian Federation presents a critical challenge to HIV control: it is the fastest growing HIV epidemic of any country [10]. Russia experienced dramatic social transformation since the collapse of the communist state with subsequent soaring rates of crime, incarceration, and poverty. These conditions, combined with a weakened healthcare infrastructure, resulted in unprecedented health crises in the 1990s, including high rates of alcohol consumption and a precipitous drop in life expectancy, largely due to excessive alcohol use [11]. Although the HIV epidemic in Russia had been initially described as attributable to intravenous drug use, [10, 12] data from 2011 show that 43.4% of new cases were due to heterosexual transmission. [13] Additionally, studies have associated alcohol consumption with HIV risk behavior [14] and suggest that certain key populations at higher risk are particularly vulnerable to the negative effects of alcohol use and HIV risk behavior, such as individuals with tuberculosis. [15] Yet the intersection between heavy alcohol use and sexual HIV risk behavior among individuals living in the RF has until recently remained largely unexplored.

One strategy to reduce alcohol-associated HIV risk is to target high-risk populations that are engaged in care; the 2008 PREVENT study of an intervention in narcology clinics in Russia is one example. [16] Given high rates of both HIV and alcohol use among TB patients, this population may represent one such important risk-group in Russia and elsewhere. Furthermore, TB treatment (requiring daily administration of medications for months) provides an opportunity to deliver interventions to reduce HIV risk behavior and alcohol consumption. For this reason, we sought to understand the complex interactions between alcohol use and HIV risk among TB patients in Russia. This study explores factors associated with HIV risk behavior in men and women with alcohol use disorders in a population of tuberculosis patients in Tomsk, Russian Federation.

Methods

Setting and population

This study was nested within a randomized controlled study, Integrated Management of Physician-delivered Alcohol Care for Tuberculosis Patients (IMPACT) in Tomsk, located in
Western Siberia (registered at clinicaltrials.gov, trial number NCT00675961). The study setting is described in detail elsewhere. [17, 18] To summarize, the IMPACT study was conducted among tuberculosis patients with alcohol use disorders (defined as any lifetime diagnosis of alcohol abuse or dependence based on the Diagnostic and Statistical Manual, 4th edition (DSM-IV) as measured by the Composite International Diagnostic Interview-Substance Abuse Module (CIDI-SAM)[19]) receiving care in a tuberculosis (TB) hospital in Tomsk Oblast, Russia, to assess the feasibility and effectiveness of integrating two modalities of alcohol treatment into the tuberculosis treatment program. [20] Tomsk Oblast has a population of 1,200,000, and, as of 2013, a tuberculosis incidence rate of 57.6 per 100,000, HIV incidence of 189.8 per 100,000 and TB/HIV prevalence of 13.4 per 100,000. [personal communication, A. Solovyova] No reliable data exist that describe alcohol use in Tomsk; however, one study found that 52% of total mortality among the 15–54 year old population in Tomsk is attributable to alcohol use. [21] Tuberculosis care in Tomsk is provided at no cost to patients and conducted under directly observed therapy. TB patients are screened for HIV and alcohol use disorders prior to initiation of TB therapy.

Ethics review and approval for the IMPACT study was provided by the Partners Healthcare Institutional Review Board; the State Research Center Virology and Biotechnology, Novosibirsk Region; and the Siberian State Medical University of the Federal Agency for Health Care and Social Development, Tomsk, Russia.

Data collection

Screening interviews were conducted on all adults starting TB treatment at the TB Hospital. The Alcohol Use Disorders Identification Test (AUDIT) [22] was routinely used by TB physicians to assess alcohol use severity [17]. Patients meeting eligibility criteria—TB patients being treated at one of the 3 study sites with a concurrent alcohol use disorder as defined by the CIDI-SAM [19]—were approached for study enrollment. After signing informed consent, study participants were randomized to one of four arms (behavioral intervention, naltrexone, both, or treatment as usual) and followed for 6 months. Baseline data on demographics, clinical history and course, alcohol use, depression and psychosocial factors were collected through interviews and chart review. Baseline depression was assessed using the Center for Epidemiologic Studies Depression (CES-D) scale [23]. Alcohol use and HIV risk behavior were assessed with the Timeline Follow-Back Calendar (TLFB)[24] and a modified Risk Assessment Behavior (RAB) tool, [25] respectively. The Addiction Severity Index Scale (ASI)[26] was also applied to assess psychosocial consequences of alcohol use.

Data analysis

The primary outcome for this analysis was baseline “HIV risk behavior,” defined as participant report of at least one of the following behaviors in the past six months: 1) high-risk injection drug use (IDU) and/or 2) multiple sexual partners with inconsistent condom use. Inconsistent condom use was defined as participant report that condoms had not been used every time the participant had sex. High-risk injection drug use was defined as injecting drugs and sharing or trading injection apparatus or supplies, or injecting drugs in the context of a shooting gallery. “Depression” was defined as a CES-D score of 16 or higher. Alcohol use severity was defined using five criteria: 1) number of drinking days (defined as a day in which any alcohol was consumed) in past month; 2) number of heavy drinking days (defined using US National Institute on Alcohol Abuse and Alcoholism standard as a day in which 4 or more standard drinks were consumed if female or 5 or more standard drinks were consumed if male) in the past month; 3) average number of drinks per drinking day; 4) average number of drinks per heavy drinking
day; and 5) “zapoi” (a Russian term for extended binge drinking), defined as any episode of at least three consecutive heavy drinking days. Social capital was assessed based on a series of questions in a baseline questionnaire addressing trust in other people and ability to turn to family or friends in time of need. A social capital score was created by totaling the responses of these questions, ranging from 0 (least trust, fewest people to turn to in time of need, least resources) to 9 (most trust in others, most people, most resources) “Low social capital” was defined as an overall social capital score less than 4.

Data were analyzed in Stata 11.2 (College Park, TX). Chi squared tests and Fisher’s exact tests were used to assess associations between HIV risk behavior and binary or categorical variables, and t-tests or Wilcoxon rank-sum tests used for continuous data. Odds ratios were calculated using single and multiple logistic regression or exact logistic regression (using conditional maximum likelihood estimates for small numbers) to assess baseline data associations. Variables significant at \( p = 0.1 \) were considered for inclusion in the multivariable analysis. Alcohol use variables were tested for collinearity before entry into the model. A final model was determined using likelihood ratio testing. Variables included in the multivariable analysis were assessed for interaction and effect modification.

Results

Two hundred patients were enrolled in the randomized controlled trial. Of these 200, four patients were withdrawn from the study when active TB was ruled out; thus, 196 participants (82.1% males) comprised the cohort for this study. Demographic and clinical data for the participants are presented by gender in Table 1. Assessment of gender differences in substance use are presented in Table 2. Of the 196 participants, 46 (23.5%) endorsed HIV risk behavior at baseline. Five (2.6%) reported only risk behavior related to IDU, 37 (18.9%) reported only sex risk, and four (2.0%) reported both. Ninety-nine (50.5%) participants reported at least one heavy drinking day in the month before starting TB treatment, and ninety-seven (49.5%) reported at least one bout of “zapoi” [data not shown]. A higher proportion of men than women reported HIV risk behavior (26.1% vs. 11.4%, borderline statistically significant at \( p = 0.06 \)). There were no gender differences in reported alcohol use including number of

| Table 1. Socio-demographic characteristics of study participants, by sex. Tomsk, Russian Federation. \( n = 196 \). |
|---|---|---|
| Factors (n, if not 196) | Total or \( n \) Mean [std] | Male, \( N = 161 \) or \( n \) [\%] | Female, \( N = 35 \) or \( n \) [\%] |
| Age (n = 195) | 41.5 [10.9] | 42 [10.9] | 39.1 [10.9] |
| Under 41 years (n = 195) | 95 (48.7) | 77 (48.1) | 18 (51.4) |
| Married or living together | 74 (37.7) | 59 (36.6) | 15 (42.9) |
| Prior incarceration | 53 (27.3) | 51 (32.8) | 2 (5.7) |
| Low social capital | 16 (8.2) | 12 (7.5) | 4 (11.4) |
| Depression | 26 (13.3) | 17 (10.6) | 9 (25.7) |
| History of sexually transmitted infection | 73 (37.2) | 58 (36.0) | 15 (42.9) |
| Current tobacco use | 182 (92.8) | 151 (93.8) | 31 (88.6) |
| Low body mass index | 56 (28.6) | 34 (21.1) | 22 (62.9) |
| Multi-drug resistant tuberculosis | 26 (13.3) | 22 (13.7) | 4 (11.4) |
| HIV positive at intake | 1 (0.5) | 0 | 1 (2.9) |

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drinking days, binge drinking episodes or drinks per drinking day. No women reported injection drug risk behavior in the six months prior to the study.

We assessed associations between demographic and clinical characteristics and HIV risk behavior at baseline (Table 3). In unadjusted analysis, having a stable partner (married or living with a partner), a low body mass index (a possible proxy for illness severity) and greater number of days of alcohol abstinence in the month prior to diagnosis were significantly associated with a lower likelihood of HIV risk behavior. Incarceration history, younger age (40 or younger), a diagnosis of drug addiction, prior history of a sexually transmitted disease, low social capital and heavier alcohol use (in the form of higher AUDIT score and high ASI score) were significantly more likely to be associated with HIV risk behavior. In adjusted analysis, age, history of prior incarceration and history of a sexually transmitted disease continued to be significantly associated with HIV risk behavior.

Discussion

This cohort of patients with tuberculosis and alcohol use disorders presents with great psychosocial vulnerabilities, including high levels of unemployment and past incarceration at the time of enrollment. Seventy-two percent of patients consecutively starting TB treatment were found to meet DSM-IV criteria for a history of alcohol abuse or dependence. Alcohol consumption in this cohort was high among both men and women, as reflected by the average ten drinks per drinking day at baseline in both genders. Although women’s alcohol consumption in Russia is known to be high (12.6 liters per capita in 2010)[27], this finding of gender parity for alcohol use in tuberculosis patients is striking and significant, as studies across many countries have found that women historically report lower levels of alcohol use than men, [28] including both WHO data [27] and a prospective study of alcohol and mortality in Russia. [29] However, these results are consistent with an earlier study in a similar population that showed that among people with AUDs in Tomsk, consumption among the women was not significantly less than that of men. [18] These high levels of alcohol consumption among female tuberculosis

| Factors (n, if not 196) | Total | Male, N = 161 Mean [std] or n (%) | Female, N = 35 Mean [std] or n (%) | P-value |
|------------------------|-------|----------------------------------|----------------------------------|---------|
| High Alcohol Severity Index (ASI) score | 22 (11.2) | 18 (11.0) | 4 (11.4) | 0.95 |
| Heavy drinking days in 30 days prior to baseline | 3.07 [5.2] | 3.07 [5.2] | 3.06 [5.5] | 0.64 |
| Mean # drinks per drinking day (n = 129) in 30 days prior to baseline | 10.3[7.99] | 10.4[7.4][8.9] | 10.1[10.4][5.4] | 0.34** |
| Mean # drinks per heavy drinking day (n = 99) in 30 days prior to baseline | 14.1[7.5] | 14.04 [6.85] | 14.2[10.32] | 0.64** |
| Number of binge drinking (zapoi) episodes (n = 193) in 30 days prior to baseline | 1.69 [2.5] | 1.61 [2.3] | 2.08 [3.1] | 0.69** |
| AUDIT score (n = 194) | 18.3 [7.7] | 18.8 [7.7] | 15.79 [7.07] | 0.05 |
| HIV risk behavior in the 6 months prior to baseline data collection | 46 (23.5%) | 42 (26.1%) | 4 (11.4%) | 0.06* |

* Fisher’s exact test  
** Wilcoxon rank sum

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patients are notable and worth greater exploration for both their clinical and public health significance.

RAB scores in general were quite low, ranging from 0 to 17 out of a possible 40, consistent with other studies among Russian TB patients. However, a high proportion (23.2%) practiced behaviors associated with high HIV risk in the six months prior to starting TB treatment, primarily the combination of inconsistent condom use (68% of the cohort) and multiple partners (26% of the cohort).

Alcohol use severity was associated with HIV risk behavior in unadjusted analysis, but lost statistical significance when adjusting for demographic and social factors. There are many possible explanations for this finding, particularly among a cohort of individuals with severe and long-standing alcohol use disorders, in which specific patterns of alcohol use may not have significantly influenced their degree of HIV risk behavior. For example, Ehrenstein et al demonstrated that in a cohort of HIV-infected people with alcohol problems, risky alcohol use (defined as >14 drinks per week in men or >7 in women) was associated with inconsistent condom use in active drug users but not in people who didn’t actively use injection drugs. More recently, Wirtz et al report that in a Russian cohort of men who have sex with men, consistent condom use during anal sex was statistically significantly lower for heavy alcohol users than low-level users. While heavy alcohol use has been associated with general increases in sexual HIV risk behavior in many settings at the global level, studies have shown a lack of association between alcohol use and condom use at the level of actual sexual event; one theory proposed is that one is either an habitual condom user or not, and alcohol does not change that behavior materially. In addition, binge drinking may pose a different risk for risky sexual behaviors than chronic alcohol use.

Table 3. Factors associated with HIV high risk behavior at baseline among study participants, Tomsk, Russian Federation (n = 196).

| Factors (n if not 196) | OR 95% CI | p-value | Adjusted OR 95% CI | P-value |
|----------------------|-----------|---------|-------------------|---------|
| Male sex*            | 2.73 (0.91, 8.21) | 0.07 | 12.11 (1.38, 106.21) | 0.02 |
| Married or living together* | 0.27 (0.12, 0.61) | <0.01 | 5.7 (0.48, 68.07) | 0.17 |
| Age less than 41 years* | 2.97 (1.46, 6.04) | <0.01 | 4.93 (2.10, 11.58) | <0.001 |
| Prior incarceration*   | 3.93 (1.95, 7.95) | <0.01 | 3.56 (1.55, 8.17) | <0.002 |
| Low social capital*   | 2.81 (0.99, 8.03) | 0.05 |
| Depression            | 1.23 (0.48, 3.15) | 0.65 |
| History of sexually transmitted infection* | 2.00 (1.02, 3.90) | 0.04 | 3.48 (1.50, 8.10) | <0.01 |
| Current tobacco use** | 0.75 (0.20, 3.45) | 0.85 |
| Low body mass index*  | 0.37 (0.15, 0.88) | 0.03 |
| Multi-drug resistant tuberculosis | 1.54 (0.62, 3.82) | 0.35 |
| Lifetime alcohol dependence disorder | 1.18 (0.59, 2.35) | 0.63 |
| Lifetime alcohol abuse disorder** | 5.01 (0.79, inf) | 0.09 |
| Lifetime drug use disorder, DSM-IV * | 3.60 (1.10, 11.77) | 0.03 |
| High Alcohol Severity Index (ASI) score* | 2.56 (1.02, 6.46) | 0.046 |
| Heavy drinking days * | 1.05 (0.99, 1.12) | 0.07 |
| Abstinent days*       | 0.96 (0.93, 0.99) | 0.03 |
| Mean # drinks per drinking day (n = 131) in 30 days prior to baseline | 1.00 (0.99, 1.02) | 0.33 |
| Mean # drinks per heavy drinking day (n = 107) in 30 days prior to baseline | 1.00 (0.99, 1.02) | 0.33 |
| AUDIT score *         | 1.05 (1.01, 1.10) | 0.02 |

*included into main effects multivariable model
**exact Logistic regression

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It is also possible that we were unable to accurately measure HIV risk behavior. While the RAB is a useful “broad brush” to risk assessment, it was initially developed for and validated specifically with injection drug users. [35] Although the RAB is quite sensitive to both drug risk and sex risks for men who have sex with men, it may be less sensitive in exclusively heterosexual men and women. The RAB asks whether condoms had been used “all of the time, most of the time, some of the time, none of the time” but does not ask with which partners condoms were used. If a person in a relationship never used condoms with his/her spouse, but used them with others, this may overestimate true risk. Other, more precise tools to assess HIV risk factors in these groups may provide better estimates of true risk.

Two main limitations of the study are the small sample size and that it is based on secondary analysis of data from a clinical trial. Our small sample size impeded our ability to explore additional gender differences in HIV risk behavior; thus, we were unable to compare factors associated with risky behavior by gender, which is known to be associated in many studies with both sexual risk [36, 37] and alcohol risk behavior. [28] Gender differences in self-report of both sex and alcohol behaviors may also exist. Some studies have documented that women face more stigma with respect to alcohol use, [38] and studies specifically in Russia have documented women underreporting alcohol use. [39] We acknowledge that findings described in this cohort of TB patients are not generalizable to the broader population; nonetheless they could inform prevention and risk reduction interventions directed at such individuals while receiving TB treatment.

As the Russian HIV epidemic continues to accelerate and the alcohol use epidemic does not show signs of slowing, understanding the complexities of HIV transmission dynamics in Russia remains an urgent priority to inform strategies to address the epidemic. Studies addressing event-level HIV risk behaviors and qualitative studies on risky behavior, particularly aimed at understanding the complex relationship between alcohol use, infectious disease, gender, and other psychosocial factors in Russia are important in designing culturally relevant intervention strategies that target specific behavior such as condom use among vulnerable populations. Larger studies, specifically addressing sex differences in risks and barriers to protective behavior, would also be important.

Supporting Information

S1 File. IMPACT Aim 3 public dataset. (CSV)

S2 File. Impact Aim 3 data dictionary. (LOG)

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Author Contributions
Conceived and designed the experiments: SS HC SFG GY SY VL. Performed the experiments: VL GY SY. Analyzed the data: ACM SA AKN CL. Wrote the paper: ACM AKN SS SFG HC. Revised and edited the manuscript: ACM AKN SS SFG SA CL VL GY SY.

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