

Alcohol consumption analysis among patients with liver disease in China

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Abstract

Background: Alcohol consumption has been observed to be a contributing factor in liver damage. However, very few studies have tried to decipher the correlation between patients with liver disease and alcohol consumption. Therefore, this study was planned to determine the prevalence of alcohol consumption among patients with liver disease, and to evaluate the risk factors, liver diseases, and chronic medical conditions associated with alcohol drinking.

Methods: A cross-sectional study was conducted among patients with liver disease in 30 provinces, autonomous regions, and municipalities across China. All participants answered the questionnaire, which led to the calculation of Alcohol Use Disorders Inventory Test (AUDIT) score for each patient. Based on this score, low-risk drinkers, hazardous drinkers, and harmful drinkers were defined as having AUDIT score of <8, between 8 and 15, and ≥ 16 , respectively.

Results: A total of 1489 participants completed the questionnaire. Based on this information, 900 (60.44%) participants were classified as alcohol drinkers. Among these, 8.66% were ex-drinkers, 22.10% were low-risk drinkers, 17.13% were hazardous drinkers, and 12.56% were harmful drinkers. Further investigation of the association between alcohol consumption and other baseline characteristics of patients with liver disease revealed that usually men <40 years old, participants having higher family annual income, having college degree or higher education, living alone, having higher body mass index (BMI), current smokers, and ex-smokers had significant association with higher risk of alcohol consumption. In addition, among the 18.07% of the participants with cirrhosis, it was observed that risk of cirrhosis increased with higher alcohol consumption. Furthermore, harmful drinkers showed greater odds of hypertension and heart diseases, while hazardous drinkers and harmful drinkers, both had greater odds of hyperlipidemia.

Conclusions: Overall our analyses indicated that among the patients with liver disease in China, there was high rate of alcohol consumption and dependence. Alcohol consumption usually associated with men <40 years old, higher family income, education level, living alone, high BMI, and smoking. Increased alcohol consumption not only increased the risk of cirrhosis, but also enhanced the risk of hypertension, heart diseases, and hyperlipidemia.

Keywords: Alcohol consumption; Patients with liver disease; Alcohol dependence; AUDIT; Chronic medical conditions

Introduction

According to the 2014 World Health Organization (WHO) report on alcohol and health, alcohol consumption has been estimated to result not only in 139 million disability-adjusted life years, but also contributes to 5.1% of the global burden of disease and injury.^[1] Typically, alcohol drinking is normal part of daily diet, especially in rural areas of China. It is also commonly consumed during important festivals, business occasions, rituals, and special

events. In this context, Chinese Health and Nutrition Survey indicated that alcohol consumption rate in the past year was around 34% in China.^[2]

The hepatitis virus infection disease burden is also high in China, and usually results in liver disease. Especially patients where liver disease stems from hepatitis B virus (HBV) or hepatitis C virus (HCV) infection, the alcohol consumption has been observed to further contribute to liver damage.^[3,4] In addition, alcohol intake has also been observed to

Access this article online

Quick Response Code:



Website:
www.cmj.org

DOI:
10.1097/CM9.000000000000067

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Chinese Medical Journal 2019;132(4)

Received: 26-11-2018 Edited by: Li-Shao Guo

independently increase the risk of cirrhosis and hepatocellular carcinoma (HCC) in patients with HBV or HCV infection.^[5] Multiple studies have indicated about the interactive correlation of heavy alcohol consumption and HBV or HCV infection, with risk of cirrhosis.^[6] Some earlier studies have even shown synergistic effect of alcohol drinking and HBV/HCV infection,^[7,8] as a mechanism of action for HCC development.^[8] Therefore, in light of high disease burden of hepatitis virus (HBV and HCV) infection, the efforts to combat liver disease development, must not only focus on primary prevention of hepatitis virus infection, but also on the other risk factors, including alcohol consumption especially in patients with hepatitis virus infection.

In parallel, alcohol consumption has also been shown to correlate with increased morbidity and mortality in certain chronic diseases.^[9-11] In this regard, a recent meta-analysis study showed 2-fold increase in cardiovascular disease and cancer mortality rate, while 15-fold increase in liver cirrhosis mortality rate in individuals with alcohol use disorder (AUD), in comparison to general population.^[12] Moreover, another study reported that despite drinking patterns, the all-cause mortality rate at 7-year follow-up was significantly higher in those individuals who met alcohol dependence criteria in the past year, relative to the general population.^[13]

An interesting study from United Kingdom revealed that general practitioners only discussed alcohol use in less than one-third of the cases that were subsequently admitted to hospital due to alcohol-related liver disease (ALD).^[14] This observation highlights an important fact that despite patients developing ALD in the community, they are not identified until they are admitted to the hospital with end stage of liver disease, and by this time, it is too late for many of them to get treatment. Similarly, the data from numerous randomized controlled trials concluded that if high-risk drinking behavior is identified early in the community or at outpatient service of hospital, then brief advice or intervention can prove to be cost-effective way to reduce drinking.^[15,16] Thus, early identification of alcohol consumption behavior should be encouraged.

Importantly, little information about alcohol consumption prevalence among Chinese patients with liver disease is available. There is also a poor understanding about the effect of alcohol consumption on liver injury and chronic medical conditions among patients with liver disease in China. Therefore, in this study, we have investigated the prevalence of alcohol drinking among patients with liver disease, and identify the association of alcohol consumption with liver disease and various chronic medical conditions in China.

Methods

Ethics approval

The process of completing questionnaires, collecting and reviewing patients' data was approved by the Institutional Review Board (IRB) of Peking University People's Hospital (IRB reference number 2018PHB038-01). All the participants provided written informed consent.

Study design

A cross-sectional study was conducted to evaluate the alcohol drinking behavior of patients with liver disease, who visited hepatology or gastroenterology outpatient department at Chinese hospitals, between March 1, 2018 and May 31, 2018. This population-based study was simultaneously conducted in 30 provinces, autonomous regions, and municipalities in China. Provincial categories included "Northern China" (Beijing, Tianjin, Hebei, Heilongjiang, Jilin, Liaoning, and Inner Mongolia), "Southern China" (Guangdong, Guangxi, Fujian, Yunnan and Hainan), "Western China" (Sichuan, Xinjiang, Chongqing, Shanxi, Shaanxi, Gansu, Ningxia, Qinghai and Guizhou), "Eastern China" (Shanghai, Shandong, Jiangxi, Jiangsu and Zhejiang), and "Central China" (Henan, Hunan, Hubei and Anhui).

The patients were excluded from the study, if they showed hepatic encephalopathy, were inebriated at the time of the interview, or had any other condition that prohibited them to properly answer the questionnaire. All the participants were interviewed face-to-face by trained investigators on the day of their regular medical appointment. All patients were explained about the goals of the study before asking them to sign the consent forms. Subsequently, these patients answered the standard general questionnaire.

Questionnaire design

Social demographics and lifestyle factors

The questionnaire included social demographic variables (age, gender, nationality, height, weight, occupation, education, family annual income, marital and living status, and area of residence) and lifestyle variables (smoking). Based on the information provided by the participants, we calculated body mass index (BMI), using height and weight ($\text{weight [kg]/height [m]}^2$). Overweight has been defined as $\text{BMI} \geq 25 \text{ kg/m}^2$ for Asian countries.^[17] In addition, smoking cessation was defined as no smoking in 3 months prior to interview.

Liver diseases

The questionnaire asked participants if they have any of the following liver diseases, diagnosed by a competent doctor; Chronic hepatitis B (no treatment or under treatment), Chronic hepatitis C (no treatment or under treatment or cured), alcoholic liver disease, non-alcoholic fatty liver disease, drug-induced liver disease, autoimmune liver disease (ie, autoimmune hepatitis, primary biliary cholangitis, primary sclerosing cholangitis), acute hepatitis A, acute hepatitis E, agnogenic abnormal liver function, liver cirrhosis, and HCC.

Chronic medical conditions

In addition questionnaire also required information from participants about the following 8 chronic medical conditions that had been confirmed by a competent doctor, hypertension, heart disease (myocardial infarction, angina pectoris, tachycardia, and other forms), diabetes mellitus, hyperlipidemia (ie, elevated of cholesterol or

triglycerides or low density lipoproteins), hyperuricemia, gout, cerebrovascular disease (ie, cerebral infarction, cerebral hemorrhage), stomach illness (ie, gastritis, duodenal, or stomach ulcer), and renal failure.

Alcohol consumption

Finally, the participants were also asked about their alcohol consumption habits. The participants reporting consumption of any alcoholic drink in the past 12 months prior to interview were classified as alcohol drinkers, while those who have no alcohol consumption at least during the same period, but had prior history were categorized as ex-drinkers. In total, our questionnaire contained eighteen different questions about alcohol consumption. Among these, questions 1 to 10 were cited from Alcohol Use Disorders Inventory Test (AUDIT), and the answers to these questions led to the calculation of AUDIT score for each patient. The questions 1 to 3 were about exploring alcohol consumption, while questions 4 to 6 were related to alcohol dependence, and questions 7 to 10 assessed alcohol-related problems. In addition, there were 2 cut-off points, 1 for dependence (AUDIT score ≥ 16) and 1 for risky drinking (AUDIT score: 8–15).^[18] In our study, low-risk drinkers were defined as those having AUDIT score of <8 , while hazardous drinkers were defined as those with AUDIT score between 8 and 15. However, those with AUDIT score of ≥ 16 were classified as harmful drinkers.

The remaining other 8 questions were “How long have you been drinking?”, “What kind of alcohol did you usually have in recent one year,” “What is the reason you have a drink in most cases,” “Over the past year, the proportion of you involuntary having a drink (ie, business entertainment),” “Have you sought professional help or been hospitalized because of adverse effects after drinking,” “Did you try to cease drinking,” “To cease drinking successfully, do you want to go to the professional clinic of abstinence,” and “Do you know that alcohol consumption is harmful to humans.”

Data collection

To collect the complete information, each patient was interviewed during his/her regular medical appointment in an outpatient department of a hospital. All patients completed the questionnaires with the assistance of an interviewer. The information about patient's baseline characteristics (age, gender, nationality, height, weight, occupation, education, family annual income, marital and living status, and area of residence), smoking history, diagnosis of liver disease and other chronic medical conditions and alcohol consumption were collected. All the patients provided complete information, and there was no missing response. The baseline data were analyzed for information about frequency and quantity of alcohol consumption and its wide-ranging associations with sociodemographic status, liver diseases, and chronic medical conditions.

Statistical analysis

The proportions of baseline characteristics among never drinkers and drinkers were compared using Chi-squared

test. Multiple logistic regression analyses were performed to estimate the odds ratios (ORs) with 95% confidence intervals (CIs), to assess association of sociodemographic and lifestyle factors, with alcohol consumption. Similar analyses were performed to assess association of liver disease and chronic medical condition among drinkers and non-drinkers. All tests were 2-sided and P -value of <0.05 represented statistical significance. All analyses were conducted using Statistical Analysis System (SAS) version 9.4 (SAS Institute, Inc., Cary, NC, USA) software.

Results

Baseline characteristics of patients with liver disease

A total of 1489 participants completed the questionnaires. Among these 615 (41.30%) were from northern China, while 96 (6.45%), 397 (26.66%), 297 (19.95%), and 95 (6.38%) patients were from southern, western, eastern, and central China, respectively. The mean age of these participants was 43 years and 1044 (70.11%) were male in gender. A total of 1396 (93.75%) participants were of Chinese Han nationality. The family annual income of 594 (39.89%) participants was less than 60,000 RMB Yuan, while 561 (37.68%) had it between 60,000 RMB Yuan and 150,000 RMB Yuan. In terms of their education status, 855 (57.42%) individuals had college or higher education degree. The 477 (32.03%) participants were white collar, and 348 (23.37%) were blue collar. In addition, 1223 (82.14%) individuals were married. Analyses of the living status revealed that 790 (53.06%) patients were living with family, while 535 (35.93%) were living alone. A total of 32.30% of the participants had BMI of ≥ 25 kg/m². The 717 (48.15%) participants were categorized as never smokers, while 300 (20.15%) were characterized as current smokers. The complete information about all the baseline characteristics of participants is summarized in Table 1.

Alcohol consumption analysis among patients with liver disease

Overall 900 (60.44%) participants were identified as alcohol drinkers, and among them, 129 (8.66%) were ex-drinkers. Alcohol drinking rate was observed to be higher among men in comparison to women ($P < 0.0001$). Among the 329 (22.10%) participants identified as low-risk drinkers, 67.17% were males and 32.83% were females. In contrast, among the 255 (17.13%) participants characterized as heavy drinkers, majority (84.91%) of them were male. Overall, a 187 (12.56%) participants met the criteria of alcohol dependence (harmful drinkers) [Table 1].

Multiple logistic regression analyses about the factors associated with alcohol consumption [Table 2], revealed that men were more likely to consume alcohol as compared to women (OR=0.274, 95% CI=0.217–0.345). In addition, patients younger than 40 years of age were more likely to drink alcohol than those aged between 50 and 59 years or aged older than 60 years (OR=0.705, 95% CI=0.528–0.940; OR=0.530, 95% CI=0.368–0.765, respectively). Higher family annual income (ranging from 60,000 to 500,000 RMB Yuan) also showed

Table 1: Baseline characteristics of participants based on alcohol consumption categories.

Variables	Total, n (%)	Never drinkers, n (%)	Ex-drinkers, n (%)	Low-risk drinkers, n (%)	Hazardous drinkers, n (%)	Harmful drinkers, n (%)	χ^2	P
Participants	1489	589 (39.56)	129 (8.66)	329 (22.10)	255 (17.13)	187 (12.56)		
Gender							176.59	<0.0001
Male	1044 (70.11)	316 (53.65)	106 (82.17)	221 (67.17)	228 (89.41)	173 (92.51)		
Female	445 (29.89)	273 (46.35)	23 (17.83)	108 (32.83)	27 (10.59)	14 (7.49)		
Age (years)							31.07	0.0019
<40	646 (43.38)	228 (38.71)	56 (43.41)	166 (50.46)	120 (47.06)	76 (40.64)		
40–49	426 (28.61)	169 (28.69)	34 (26.36)	89 (27.05)	81 (31.76)	53 (28.34)		
50–59	275 (18.47)	120 (20.37)	20 (15.50)	53 (16.11)	39 (15.29)	43 (22.99)		
≥60	142 (9.54)	72 (12.22)	19 (14.73)	21 (6.38)	15 (5.88)	15 (8.02)		
Nationality							12.00	0.0174
Han	1396 (93.75)	561 (95.25)	120 (93.02)	313 (95.14)	236 (92.55)	166 (88.77)		
Non-Han	93 (6.25)	28 (4.75)	9 (6.98)	16 (4.86)	19 (7.45)	21 (11.23)		
Family annual income (RMB Yuan)							76.11	<0.0001
<60,000	594 (39.89)	290 (49.24)	60 (46.51)	112 (34.04)	72 (28.24)	60 (32.09)		
60,000–149,999	561 (37.68)	203 (34.47)	49 (37.98)	134 (40.73)	101 (39.61)	74 (39.57)		
150,000–299,999	241 (16.19)	74 (12.56)	15 (11.63)	58 (17.63)	52 (20.39)	42 (22.46)		
300,000–500,000	54 (3.63)	11 (1.87)	3 (2.33)	15 (4.56)	22 (8.63)	3 (1.60)		
>500,000	39 (2.62)	11 (1.87)	2 (1.55)	10 (3.04)	8 (3.14)	8 (4.28)		
Education level							81.43	<0.0001
Below middle school	62 (4.12)	34 (5.77)	8 (6.20)	14 (4.26)	1 (0.39)	5 (2.67)		
Middle school or high school	572 (38.42)	284 (48.22)	50 (38.76)	83 (25.23)	75 (29.41)	80 (42.78)		
College and above	855 (67.42)	271 (46.01)	71 (55.04)	232 (70.52)	179 (70.20)	102 (54.55)		
Occupation							29.97	0.0002
White collar	477 (32.03)	180 (30.56)	35 (27.13)	128 (38.91)	82 (32.16)	52 (27.81)		
Blue collar	348 (23.37)	122 (20.71)	34 (26.36)	70 (21.28)	55 (21.57)	67 (35.83)		
Others	664 (44.59)	287 (48.73)	60 (46.51)	131 (39.82)	118 (46.27)	68 (36.36)		
Marital status							18.92	0.0153
Married	1223 (82.14)	482 (81.83)	96 (74.42)	276 (83.89)	217 (85.10)	152 (81.28)		
Single	177 (11.89)	74 (12.56)	22 (17.05)	40 (12.16)	26 (10.20)	15 (8.02)		
Others	89 (5.98)	33 (5.60)	11 (8.53)	13 (3.95)	12 (4.71)	20 (10.70)		
Living status							19.84	0.0109
Living with family	790 (53.06)	340 (57.72)	58 (44.96)	171 (51.98)	130 (50.98)	91 (48.66)		
Living with others	164 (11.01)	58 (9.85)	25 (19.38)	33 (10.03)	23 (9.02)	25 (13.37)		
Living alone	535 (35.93)	191 (32.43)	46 (35.66)	125 (37.99)	102 (40.00)	71 (37.97)		
Body mass index (kg/m ²)							66.55	<0.0001
<18.5	59 (3.96)	42 (7.13)	1 (0.78)	7 (2.13)	5 (1.96)	4 (2.14)		
18.5–24.9	949 (63.73)	412 (69.95)	81 (62.79)	213 (64.74)	140 (54.90)	103 (55.08)		
≥25	481 (32.30)	135 (22.92)	47 (36.43)	109 (33.13)	110 (43.14)	80 (42.78)		
Smoking							242.98	<0.0001
Never smoker	717 (48.15)	399 (67.74)	42 (32.56)	162 (49.24)	81 (31.76)	33 (17.65)		
Current smoker	300 (20.15)	102 (17.32)	45 (34.88)	60 (18.24)	52 (20.39)	41 (21.93)		
Ex-smoker	472 (31.70)	88 (14.94)	42 (32.56)	107 (32.52)	122 (47.84)	113 (60.43)		

significant association with higher risk of alcohol consumption (OR=1.682, 95% CI=1.329–2.129; OR=2.153, 95% CI=1.567–2.957; OR=3.729, 95% CI=1.886–7.372, OR=2.428, 95% CI=1.187–4.967, respectively). Moreover, patients who have completed college or have higher education were also observed to consume higher amount of alcohol than those who had only completed middle school education (OR=2.617, 95% CI=1.555–4.403). The analysis of the living status revealed that patients living alone were more likely to consume alcohol than those living with family (OR=1.361, 95% CI=1.085–1.706). Importantly, increased BMI also showed correlation with higher risk of alcohol consumption (OR=3.220, 95% CI=1.807–5.739; OR=6.330, 95% CI=3.483–11.505, for BMI 18.5–24.9 and >25, respectively). The smoking status also showed interesting correlation, as current and ex-smokers were observed to more likely consume alcohol than never smokers (OR=2.436, 95% CI=1.840–3.225; OR=5.475, 95% CI=4.161–7.205, respectively). However, no significant relationship was found between nationality or area of residence or occupation or marital status with alcohol consumption.

The data from AUDIT analysis are tabulated in Figure 1. A total of 43.56% of the alcohol drinking population consumed alcohol drink monthly or less, while 54.22% of them had 1 or 2 drinks on a typical day. Very small proportion (3.78%) of the patients had a drink containing alcohol 4 or more times a week, and 10 or more drinks each time [Figure 1A]. In addition, it was also noticed that 41.12% of the participants had 5 or more drinks on 1 occasion monthly. The 45.22% of them reported that they were not able to stop drinking once they had started at least monthly during the last year, and 41.44% failed to respond normally after drinking during the same time period. The 26.33% of them realized that they require a drink early in the morning to get themselves going after a heavy drinking session, at least monthly during the last year. The 44.33% of them had a feeling of guilt or remorse after drinking, while 40.89% were unable to remember what happened the night before because of their monthly drinking habit during the last year [Figure 1B]. The 21.33% reported that either they or someone else was injured due to their alcohol drinking. Interestingly, 58.56% of the participants revealed that either their relative, friend, doctor, or other health care worker were

Table 2: Multiple logistic regression analyses of factors associated with alcohol consumption.

Variables	Never drinkers, n (%)	Drinkers, n (%)	Odds ratio	95% CI	P
Participants	589 (39.56)	900 (60.44)			
Gender					
Male	316 (53.65)	728 (80.89)	1		
Female	273 (46.35)	172 (19.11)	0.274	0.217–0.345	<0.0001
Age (years)					
<40	228 (38.71)	418 (46.44)	1		
40–49	169 (28.69)	257 (28.56)	0.829	0.644–1.068	0.1466
50–59	120 (20.37)	155 (17.22)	0.705	0.528–0.940	0.0171
≥60	72 (12.22)	70 (7.78)	0.530	0.368–0.765	0.0007
Nationality					
Han	561 (95.25)	835 (92.78)	1		
Non-Han	28 (4.75)	65 (7.22)	1.560	0.989–2.460	0.0560
Family annual income (RMB Yuan)					
<60,000	290 (49.24)	304 (33.78)	1		
60,000–149,999	203 (34.47)	358 (39.78)	1.682	1.329–2.129	<0.0001
150,000–299,999	74 (12.56)	167 (18.56)	2.153	1.567–2.957	<0.0001
300,000–500,000	11 (1.87)	43 (4.78)	3.729	1.886–7.372	0.0002
>500,000	11 (1.87)	28 (3.11)	2.428	1.187–4.967	0.0151
Education level					
Below middle school	34 (5.77)	28 (3.11)	1		
Middle school or high school	284 (48.22)	288 (32.00)	1.231	0.727–2.084	0.4384
College and above	271 (46.01)	584 (64.89)	2.617	1.555–4.403	0.0003
Occupation					
White collar	180 (30.56)	297 (33.00)	1		
Blue collar	122 (20.71)	226 (25.11)	1.123	0.842–1.497	0.4304
Others	287 (48.73)	377 (41.89)	0.796	0.626–1.013	0.0632
Marital status					
Married	482 (81.83)	741 (82.33)	1		
Single	74 (12.56)	103 (11.44)	0.905	0.657–1.247	0.5426
Others	33 (5.60)	56 (6.22)	1.104	0.707–1.723	0.6636
Living status					
Living with family	340 (57.72)	450 (50.00)	1		
Living with others	58 (9.85)	106 (11.78)	1.381	0.973–1.959	0.0705
Living alone	191 (32.43)	344 (38.22)	1.361	1.085–1.706	0.0076
Body mass index (kg/m ²)					
<18.5	42 (7.13)	17 (1.89)	1		
18.5–24.9	412 (69.95)	537 (59.67)	3.220	1.807–5.739	<0.0001
≥25	135 (22.92)	346 (38.44)	6.330	3.483–11.505	<0.0001
Smoking					
Never smoker	399 (67.74)	318 (35.33)	1		
Current smoker	102 (17.32)	198 (22.00)	2.436	1.840–3.225	<0.0001
Ex-smoker	88 (14.94)	384 (42.67)	5.475	4.161–7.205	<0.0001

concerned about their drinking pattern and recommended to reduce alcohol consumption [Figure 1C].

Overall, total of 218 (24.22%) participants reported that they have been consuming alcohol from 1 to 5 years, while 209 (23.22%) have been doing it between 6 and 10 years. Among the type of drinks mostly consumed among these participants were white wine (50.01%) and beer (32.78%). Four hundred thirty (47.78%) patients mentioned social contact as the main reason for their alcohol consumption. A total of 519 (57.67%) participants never tried to stop drinking, while 252 (28.00%) of them tried but failed. Further, to successfully stop drinking, 387 (43.00%) patients wanted to go to the

professional clinic of abstinence. However, 290 (32.22%) patients believed that moderate alcohol consumption was not harmful.

Correlation between alcohol consumption and liver diseases

We further analyzed the correlation between alcohol consumption and liver disease in these patients. It was observed that fair number of patients were primarily infected with HBV (691/1489, 46.41%). Notably, among the 269 (18.07%) participants having cirrhosis, its proportion was significantly high in harmful drinkers ($P < 0.001$). On the contrary, we observed no significant difference between drinkers and never drinkers in the rate of HCC among 36 (2.42%) patients with HCC [Table 3].

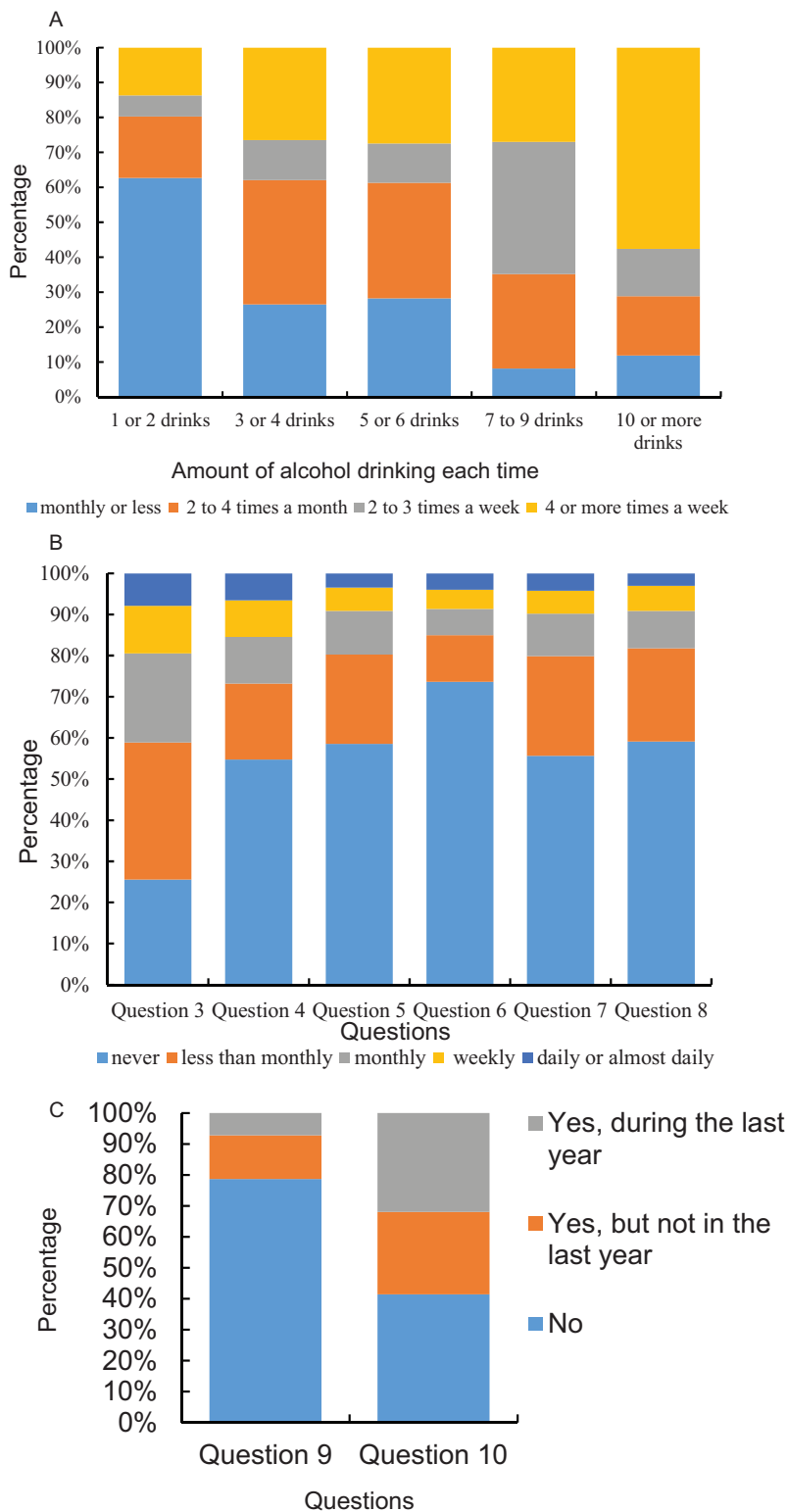


Figure 1: Tabulation of the results from Alcohol Use Disorders Inventory Test (AUDIT). (A) Question 1: How often do you have a drink containing alcohol? Question 2: How many drinks containing alcohol do you have on a typical day when you are drinking? (B) Question 3: How often do you have 5 or more drinks on one occasion? Question 4: How often during the last year you found that you were not able to stop drinking once you had started? Question 5: How often during the last year have you failed to do what was normally expected of you because of drinking? Question 6: How often during the last year you required a first drink in the morning to get yourself going after a heavy drinking session? Question 7: How often during the last year have you had a feeling of guilt or remorse after drinking? Question 8: How often during the last year have you been unable to remember what happened the night before because of your drinking? (C) Question 9: Have you or someone else been injured because of your drinking? Question 10: Has a relative, friend, doctor, or other health care worker been concerned about your drinking or suggested you to cut down?

Table 3: Prevalence of liver diseases and chronic medical conditions among patients categorized based on alcohol consumption, n (%).

Variables	Never drinkers (n=589)	Ex-drinkers (n=129)	Low-risk drinkers (n=329)	Hazardous drinkers (n=255)	Harmful drinkers (n=187)	χ^2	P
Liver diseases						149.51	<0.0001
HBV	360 (61.12)	78 (60.47)	117 (35.56)	78 (30.59)	58 (31.02)		
HCV	31 (5.26)	5 (3.88)	10 (3.04)	25 (9.80)	17 (9.09)		
ALD	0 (0)	17 (13.18)	24 (7.29)	72 (28.24)	81 (43.32)		
NAFLD	42 (7.13)	6 (4.65)	52 (15.81)	0 (0)	0 (0)		
Others*	156 (26.49)	23 (17.83)	126 (38.30)	80 (31.37)	31 (16.58)		
Liver cirrhosis	140 (23.77)	28 (21.71)	30 (9.12)	23 (9.02)	48 (25.67)	53.29	<0.0001
HBV	118 (84.29)	15 (53.57)	23 (76.67)	9 (39.13)	17 (35.42)		
HCV	6 (4.29)	1 (3.57)	0 (0)	2 (8.70)	5 (10.42)		
ALD	0 (0)	10 (35.71)	1 (3.33)	9 (39.13)	25 (52.08)		
NAFLD	6 (4.29)	0 (0)	3 (10.00)	0 (0)	0 (0)		
Others*	10 (7.14)	2 (7.14)	3 (10.00)	3 (1.18)	1 (2.08)		
HCC	16 (2.72)	1 (0.78)	9 (2.74)	4 (1.57)	6 (3.21)	3.11	0.5390
Hypertension	76 (12.90)	22 (17.05)	36 (10.94)	54 (21.18)	62 (33.16)	53.79	<0.0001
Heart disease	20 (3.40)	10 (7.75)	13 (3.95)	12 (4.71)	27 (14.44)	36.81	<0.0001
Diabetes mellitus	43 (7.30)	12 (9.30)	18 (5.47)	21 (8.24)	28 (14.97)	15.50	0.0038
Hyperlipidemia	44 (7.47)	13 (10.08)	43 (13.07)	53 (20.78)	52 (27.81)	62.91	<0.0001
Hyperuricemia	4 (0.68)	2 (1.55)	10 (3.04)	11 (4.31)	12 (6.42)	22.94	0.0001
Gout	19 (3.23)	2 (1.55)	13 (3.95)	22 (8.63)	13 (6.95)	17.24	0.0017
Cerebrovascular disease	10 (1.70)	3 (2.33)	2 (0.61)	4 (1.57)	9 (4.81)	11.98	0.0250
Stomach illness	76 (12.90)	26 (20.16)	50 (15.20)	39 (15.29)	27 (14.44)	4.74	0.3154
Renal failure	10 (1.70)	2 (1.55)	4 (1.22)	2 (0.78)	9 (4.81)	11.72	0.0451

ALD: Alcoholic liver disease, HBV: Hepatitis B virus, HCV: Hepatitis C virus, NAFLD: Non-alcoholic liver disease, HCC: Hepatocellular carcinoma.
* Drug-induced liver disease, autoimmune liver disease, acute hepatitis A, acute hepatitis E, agnogenic abnormal liver function.

Overall multiple logistic-regression-based analyses of association between alcohol consumption and liver diseases [Table 4], indicated that after adjustment, risk of alcoholic liver disease increased with increase in alcohol consumption (for low-risk drinkers, OR=4.480, 95% CI=2.087–9.616; for hazardous risk drinkers, OR=15.838, 95% CI=7.645–32.814; for harmful drinkers, OR=37.977, 95% CI=18.252–79.020, respectively). In addition, low-risk drinkers also displayed greater odds of NAFLD compared to never drinkers (OR=2.536, 95% CI=1.563–4.114). The risk of liver cirrhosis also increased with increase of alcohol consumption (for low-risk drinkers, OR=1.358, 95% CI=1.228–2.561; for hazardous risk drinkers, OR=3.337, 95% CI=2.202–4.562; for harmful drinkers, OR=3.983, 95% CI=2.630–4.533, respectively).

Association of alcohol consumption and chronic medical conditions

Typically, chronic medical conditions were quite common among men and women who participated in our study [Table 3]. Thus, we also analyzed the association between alcohol consumption and chronic medical conditions using multiple logistic regression analyses [Table 5]. It is important to note that we excluded several health conditions associated with alcohol consumption due to few number of cases for each of them like hyperuricemia, gout, cerebrovascular disease, and renal failure. However, after adjustment for variables, it was observed that harmful drinkers had greater odds of hypertension and

heart diseases in comparison to never drinkers (OR=2.443, 95% CI=1.539–3.879; OR=6.009, 95% CI=2.839–12.716, respectively). In addition, hazardous and harmful drinkers also had greater odds of hyperlipidemia, as compared to never drinkers (OR=2.468, 95% CI=1.517–4.016; OR=3.786, 95% CI=2.288–6.265, respectively). However, no significant correlation was recorded between diabetes mellitus or stomach illness and alcohol consumption.

Discussion

There have been few detailed studies focused on analyzing the alcohol consumption among patients with liver disease in China. The major highlights of our study have been (1) among patients with liver disease, the alcohol drinking rate, and alcohol dependence (harmful drinking) rate in China were 60.44% and 12.56%, respectively; (2) alcohol consumption typically showed association with men younger than 40 years old, those having high family annual income, had college or higher education, living alone and higher BMI; (3) liver cirrhosis risk showed increased trend with increase in alcohol consumption; and (4) alcohol dependence (harmful drinking) enhanced the risk of hypertension, heart disease, and hyperlipidemia.

Importantly, our study noticed a higher prevalence of alcohol consumption than previously published studies,^[2,19,20] and this could probably be related to the loose definition of “alcohol drinker.” However, it seems that loose definition was necessary to maximize the identifica-

Table 4: Multiple logistic regression analyses of factors associated with liver diseases.

Liver diseases	Alcohol consumption	n/N (%)	Unadjusted		Adjusted	
			OR (95% CI)	P	OR (95% CI)	P
HBV	Never drinkers	360/589 (61.12)	1	–	1	–
	Ex-drinker	78/129 (60.47)	0.973 (0.659–1.437)	0.8901	0.819 (0.541–1.241)	0.3467
	Low-risk drinkers	117/329 (35.56)	0.351 (0.265–0.464)	<0.0001	0.329 (0.243–0.447)	<0.0001
	Hazardous drinkers	63/255 (24.71)	0.209 (0.150–0.290)	<0.0001	0.166 (0.115–0.240)	<0.0001
	Harmful drinkers	53/187 (28.34)	0.252 (0.176–0.360)	<0.0001	0.184 (0.123–0.275)	<0.0001
HCV	Never drinkers	31/589 (5.26)	1	–	1	–
	Ex-drinker	5/129 (3.88)	0.726 (0.277–1.904)	0.5149	0.724 (0.268–1.959)	0.5248
	Low-risk drinkers	10/329 (3.04)	0.564 (0.273–1.166)	0.1224	0.599 (0.281–1.278)	0.1849
	Hazardous drinkers	10/255 (3.92)	0.735 (0.355–1.522)	0.4068	0.805 (0.360–1.801)	0.5982
	Harmful drinkers	12/187 (6.42)	1.234 (0.621–2.455)	0.5485	1.262 (0.576–2.765)	0.5608
ALD	Ex-drinker	17/129 (13.18)	1	–	1	–
	Low-risk drinkers	24/329 (7.29)	4.556 (2.151–9.651)	<0.0001	4.480 (2.087–9.616)	0.0001
	Hazardous drinkers	72/255 (28.24)	16.668 (8.351–33.267)	<0.0001	15.838 (7.645–32.814)	<0.0001
	Harmful drinkers	81/187 (43.32)	39.643 (19.886–79.028)	<0.0001	37.977 (18.252–79.020)	<0.0001
NAFLD	Never drinkers	32/589 (5.43)	1	–	1	–
	Ex-drinker	6/129 (4.65)	0.849 (0.348–2.075)	0.7204	0.771 (0.309–1.926)	0.5776
	Low-risk drinkers	52/329 (15.81)	3.267 (2.056–5.193)	<0.0001	2.536 (1.563–4.114)	0.0002
Liver cirrhosis	Never drinkers	140/589 (23.77)	1	–	1	–
	Ex-drinker	28/129 (21.71)	0.889 (0.562–1.408)	0.6162	0.799 (0.484–1.318)	0.3789
	Low-risk drinkers	30/329 (9.12)	1.322 (1.211–2.490)	<0.0001	1.358 (1.228–2.561)	<0.0001
	Hazardous drinkers	23/255 (9.02)	3.318 (2.199–4.508)	<0.0001	3.337 (2.202–4.562)	<0.0001
	Harmful drinkers	48/187 (25.67)	3.108 (2.758–4.618)	<0.0001	3.983 (2.630–4.533)	<0.0001

ALD: Alcoholic liver disease; CI: Confidence interval; HBV: Hepatitis B virus; HCV: Hepatitis C virus; NAFLD: Non-alcoholic liver disease; OR: Odds ratio.

tion of all alcohol drinkers among patients with liver disease. More specifically, it was noticed that men had higher prevalence of alcohol consumption than women in China, and this finding was consistent with previous studies^[1,19,20] and can be attributed to cultural values and norms.^[21] Typically, Chinese men has a long history of alcohol consumption, in comparison to women. The proportion of female low-risk drinkers was 32.83%. As studies have demonstrated that women are more susceptible toward hepatotoxic effects of alcohol, it is advised that women should avoid alcohol drinking.

Our analysis also indicated that both current smokers and ex-smokers were more likely to consume alcohol in comparison to never smokers and was consistent with other published studies showing a direct relationship between alcohol consumption and smoking.^[2,11,22,23] It has been reported that smoking exacerbates the effects of alcohol in inducing severe liver injury and favors development of HCC among patients with liver disease.^[22,23]

Earlier studies have also indicated about the relationship between alcohol consumption and BMI,^[24,25] and we also observed that alcohol consumption was associated with higher BMI. It has been shown that obesity is one of the most important environmental risk factor determining the risk of cirrhosis in heavy drinkers,^[26] and heavy drinkers who are overweight for at least 10 years usually have a 2-fold risk of developing cirrhosis.

The association between educational level and alcohol drinking has also been explored earlier and some studies

have suggested that less education directly correlates to alcohol dependence,^[27,28] while other studies showed that education is related to increased daily alcohol consumption or problematic drinking.^[29,30] Even some studies have indicated about no correlation between these 2 factors.^[31] However, we in our study observed that patients with higher educational levels were more likely to drink alcohol.

In terms of living status and alcohol consumption, the study conducted in Norwegian women and men showed that living with a spouse or partner was positively associated with alcohol drinking.^[32] However, we did not observe similar significant association in China. On the contrary, we noticed that living alone was more linked to higher alcohol consumption than living with family. These differences can be attributed to different drinking cultures. It has been observed that Chinese people drink more frequently at social occasions than with their spouse or partner at home. Another study showed that income levels were weakly associated with risk of heavy drinking.^[30] However, our study revealed that increase in family annual income was significantly associated with higher risk of alcohol consumption.

Generally, NAFLD diagnosis requires exclusion of daily alcohol consumption by ≥ 30 g for men and 20g for women.^[33] However, the relationship between alcohol and liver injury depends on several cofactors like alcohol type, duration of exposure, drinking patterns, and individual susceptibility, thus rendering simple quantitative thresholds partly arbitrary. Specifically, patients with moderate alcohol drinking may still be predisposed to NAFLD, if they have other metabolic risk factors.^[34] In our study, we

Table 5: Multiple logistic regression analyses of factors associated with chronic medical conditions.

Comorbidities	Alcohol consumption	n/N (%)	Unadjusted		Adjusted	
			OR (95% CI)	P	OR (95% CI)	P
Hypertension	Never drinkers	76/589 (12.90)	1	–	1	–
	Ex-drinker	22/129 (17.05)	1.388 (0.827–2.330)	0.2151	0.973 (0.547–1.730)	0.9252
	Low-risk drinkers	36/329 (10.94)	0.829 (0.544–1.264)	0.3845	0.750 (0.472–1.191)	0.2224
	Hazardous drinkers	54/255 (21.18)	1.813 (1.234–2.665)	0.0024	1.476 (0.940–2.319)	0.0910
	Harmful drinkers	62/187 (33.16)	3.348 (2.271–4.936)	<0.0001	2.443 (1.539–3.879)	<0.0001
Heart disease	Never drinkers	20/589 (3.40)	1	–	1	–
	Ex-drinker	10/129 (7.75)	2.391 (1.091–5.238)	0.0294	2.329 (0.985–5.506)	0.0540
	Low-risk drinkers	13/329 (3.95)	1.170 (0.574–2.385)	0.6648	1.571 (0.731–3.376)	0.2468
	Hazardous drinkers	12/255 (4.71)	1.405 (0.676–2.919)	0.3622	1.968 (0.854–4.532)	0.1118
	Harmful drinkers	27/187 (14.44)	4.801 (2.624–8.785)	<0.0001	6.009 (2.839–12.716)	<0.0001
Diabetes mellitus	Never drinkers	43/589 (7.30)	1	–	1	–
	Ex-drinker	12/129 (9.30)	1.302 (0.666–2.546)	0.4399	1.003 (0.482–2.089)	0.9936
	Low-risk drinkers	18/329 (5.47)	0.735 (0.417–1.296)	0.2875	0.823 (0.446–1.518)	0.5325
	Hazardous drinkers	21/255 (8.24)	1.140 (0.662–1.963)	0.6378	1.192 (0.641–2.215)	0.5784
	Harmful drinkers	28/187 (14.97)	2.236 (1.346–3.715)	0.0019	1.802 (0.983–3.303)	0.0569
Hyperlipidemia	Never drinkers	44/589 (7.47)	1	–	1	–
	Ex-drinker	13/129 (10.08)	1.388 (0.724–2.660)	0.3230	1.205 (0.612–2.376)	0.5895
	Low-risk drinkers	43/329 (13.07)	1.862 (1.195–2.903)	0.0061	1.595 (0.998–2.550)	0.0512
	Hazardous drinkers	53/255 (20.78)	3.250 (2.112–5.001)	<0.0001	2.468 (1.517–4.016)	<0.0001
	Harmful drinkers	52/187 (27.81)	4.771 (3.062–7.434)	<0.0001	3.786 (2.288–6.265)	<0.0001
Stomach illness	Never drinkers	76/589 (12.90)	1	–	1	–
	Ex-drinker	26/129 (20.16)	1.704 (1.041–2.790)	0.0341	1.772 (1.061–2.958)	0.0287
	Low-risk drinkers	50/329 (15.20)	1.210 (0.823–1.779)	0.3332	1.258 (0.841–1.884)	0.2643
	Hazardous drinkers	39/255 (15.29)	1.219 (0.803–1.850)	0.3531	1.324 (0.836–2.096)	0.2321
	Harmful drinkers	27/187 (14.44)	1.139 (0.709–1.829)	0.5900	1.160 (0.692–1.946)	0.5733

CI: Confidence interval; OR: Odds ratio.

found that low-risk drinkers had greater odds of developing NAFLD compared to never drinkers, thereby suggesting that it is better not to drink at all to avoid NAFLD.

Furthermore, we also identified that increased alcohol consumption led to increased risk of cirrhosis. It is very clear that there is a direct dose relationship between the amount of alcohol intake and likelihood of liver injury; yet, extensive variability exists between individuals. Only about 10% to 20% of the individuals with chronic heavy alcohol consumption usually develop advanced liver disease and cirrhosis, thereby indicating that additional disease modifiers and cofactors, such as behavioral, environmental, and genetic factors possibly play an important role.^[35] The participants in our study commonly consumed white wine and beer, which seems to be consistent with local lifestyle. However due to small number of cases, the alcohol type effect could not be analyzed for the risk of cirrhosis and HCC. Previous studies also could not assess the impact of alcohol type (wine *vs.* beer *vs.* liquors) and drinking patterns (ie, binge drinking and drinking outside meals) on cirrhosis and HCC,^[35] thus indicating toward large epidemiological studies.

Interestingly, alcohol consumption leads to social, family, occupational, and psychologic damage, as along with clinical comorbidities. Multiple studies have shown higher risk of hypertension,^[24] cardiovascular disease,^[36] and

heperlipemia,^[37] due to increased alcohol intake. In the present study, we also observed that alcohol dependence (harmful drinking) increased the risk of important chronic medical conditions (hypertension, heart diseases, and hyperlipidemia). However, causal relationship between alcohol consumption and chronic medical conditions could not be examined due to the cross-sectional nature of our study. But, after statistical adjustment for social-demographic factors, we indeed described the persisted association of alcohol intake with chronic medical conditions. Consistent with our data, results of another study also suggested that elevated rates of hypertension and hyperlipidemia among harmful drinkers might increase mortality rate due to coronary heart disease or cerebrovascular disease.^[38] Therefore, it is always advised to reduce alcohol drinking so as to lower the risk of hypertension, heart diseases, and hyperlipidemia.

Finally, it is important to highlight some of the limitations and important implications of our study. First, cross-sectional design precluded our study from examining causal relationships, thereby indicating toward additional randomized controlled or cohort studies. Second, the information about sociodemographic status, alcohol consumption, liver diseases and chronic medical conditions was self-reported, thus more prone to information bias and/or recall bias. However, some studies have shown that self-reported information was reasonably reproducible, suggesting reliability.^[39,40] Third, our study lacked precise quantification of alcohol consumption. While self-

reported information/categories could be imperfect, but they do identify harmful drinkers, and our outcome data support that. Moreover, our current analytic categories of “low-risk drinkers,” “hazardous drinkers,” and “harmful drinkers” were based on the AUDIT score, a widely used internationally validated WHO tool. Fourth, participants were not chosen to directly represent Chinese population as they being younger and better educated, but it still represents a group of Chinese people, who are community embedded, and generally reflective of the Chinese population, in terms of other social and economic characteristics.

Conclusion

In summary, our study predicted that prevalence of alcohol consumption and alcohol dependence among patients with liver disease was high in China. In addition, patients with liver disease commonly acknowledged that alcohol drinking is harmful, which indicated that these individuals still drank heavily despite knowing the consequences. Thus, it is imperative to devise new strategies to raise public awareness about the harmful effects of alcohol, screen alcohol drinking, and conduct brief intervention sessions in the outpatient department. It would also be helpful to set up abstinence clinics or organizations, with intent to convince patients with liver disease to stay away from alcohol consumption.

Acknowledgements

A total of 39 hospitals in 30 provinces, autonomous regions, and municipalities across China were involved. The authors thank all the investigators of these hospitals who conducted this survey.

Conflicts of interest

None.

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How to cite this article: Xie YD, Zhao CQ, Wang JP, Cheng CS, Zhou JY, Zhang Y, Xia H, Wei L. Alcohol consumption analysis among patients with liver disease in China. *Chin Med J* 2019;132:420–430. doi: 10.1097/CM9.0000000000000043