Betel nut chewing as a risk factor for hepatitis C infection in Taiwan - A Community-Based Study

To the Editor: We completed the comprehensive health survey and structural questionnaires in the adult living in the east area of Taichung City, Taiwan, including the anti-hepatitis C (HCV) antibody status and the components of metabolic syndrome from 2004 to 2005. A total of 1045 people were enrolled in this study. The anti-HCV seropositive (anti-HCV+) prevalence was 6.6% (69/1045). Results were expressed as mean±standard deviation. A *p* value < 0.05 was considered statistically significant. Multivariate logistic regression analyses were conducted to explore the factors that were independently associated with anti-HCV. The strength of association was presented as odds ratio (OR) with 95% confidence intervals (CI) and *p* values.

Table 1 shows the basic characteristics of all subjects. The anti-HCV+ subjects had higher alanine aminotransferase (ALT) and aspartate aminotransferase (AST) levels, and were elder than the anti-HCV- subjects. In addition, triglycerides levels were significantly lower in the anti-HCV+ subjects compared with the anti-HCV- subjects (p<0.01). There were no significant differences between anti-HCV+ and anti-HCV- subjects in terms of gender, body mass index (BMI), high-density lipoprotein cholesterol (HDL-C) levels, systolic and diastolic blood pressure, and fasting plasma glucose.

Multivariate logistic regression analyses were conducted to clarify the independent factors associated with anti-HCV. Variables included age, sex, smoking, drinking, betel nut chewing, exercise, milk drinking, and the presence of hypertension, diabetes mellitus, and hyperlipidemia. Betel nut chewing was significantly associated with anti-HCV+ (OR, 7.55; 95% CI, 2.51–22.74; p <0.01) as well as with milk drinking (OR, 1.63; 95% CI, 1.00–2.65; p =0.05). After adjusted for appropriated covariates, betel nut chewing was still significantly associated with anti-HCV+ (OR, 9.12; 95% CI, 2.96–28.18; p <0.01) (Table 2).

Chronic HCV infections are the major etiologies of hepatocellular carcinoma HCC in Taiwan [1]. The prevalence of anti-HCV (6.6%) in our study was higher than those in community controls [2]. The prevalence

variation in crude HCV seroprevalence ranged from 0.4 to 10.5% and HCV infection takes 2–4 decades to lead to HCC [3]. The habit of betel nut chewing is prevalent among men in Taiwan. It is especially prevalent among blue-collar workers, less educated men, cigarette smokers, and alcohol drinkers [4]. The population of betel nut chewers increased gradually. Recently, the habit of betel nut chewing was found to be a risk factor of hepatocellular carcinoma (HCC), and an increased HCC risk is associated with seropositivity of anti-HCV in Taiwan [5]. This information indirectly supports our finding that betel nut chewing is an independent risk factor of anti-HCV. Chronic hepatitis C and betel nut chewing is still a major public health concern in Taiwan. Although the precise mechanism for the association between betel nut chewing and anti-HCV remains to be determined, this study suggests that abstention from betel nut chewing is important for the prevention of chronic hepatitis C subjects.

## References

- 1. Raza SA, Clifford GM, Franceschi S. Worldwide variation in the relative importance of hepatitis B and hepatitis C viruses in hepatocellular carcinoma: a systematic review. Br J Cancer 2007;96:1127–1134
- 2. Tsai JF, Jeng JE, Ho MS, Chang WY, Lin ZY, Tasi JH. Independent and additive effect modification of hepatitis C and B viruses infection on the development of chronic hepatitis. J Hepatol 1996;**24**: 271–276.3
- Tsai MC, Kee KM, Chen YD, Lin LC, Tasi LS, Chen HH, Lu SN. Excess mortality of hepatocellular carcinoma and morbidity of liver cirrhosis and hepatitis in HCV-endemic areas in an HBV-endemic country: geographic variations among 502 villages in southern Taiwan. J Gastroenterol Hepatol 2007; 22: 92–98
- 4. Chew JW, Shaw JH. A study on betel quid chewing behavior among Kaohsiung residents aged 15 years and above. J Oral Pathol Med 1996; 25: 140–143.
- 5. Liu CJ, Chen CL, Chang KW, Chu CH,Liu TY. Safrole in betel quid may be a risk factor for hepatocellular carcinoma: case report. Can Med Assoc J 2000;162: 359–360.27.

Table1. Basic characteristics of anti-HCV seropositive and negative subjects*					
	Case	Control	p-value		
	Postive n=69(%)	Negative n=976(%)			
Male	26(37.7)	365(37.4)	0.96		
Female	43(62.3)	611(62.6)			
Age(year)	60.91±11.55	55.94±14.04	< 0.01		
Height(cm)	157.42±7.51	159.11±7.69	0.08		
Waist(cm)	62.16±8.35	63.01±11.27	0.43		
BMI(kg/m2)	25.15±3.96	24.84±3.69	0.54		
Systolic pressure(mmHg)	133.26±22.08	133.10±20.93	0.95		
Diastolic pressure(mmHg)	81.26±13.77	80.92±13.67	0.84		
AST(IU/L)	38.55±27.37	24.56±37.52	< 0.01		
ALT(IU/L)	40.19±37.30	23.31±49.91	< 0.01		
Triglycerides(mmol/dL)	$1.14 \pm 0.58$	1.43±0.98	< 0.01		
FPG(mmol/L)	5.47±1.64	5.73±2.05	0.29		
HDL-cholesterol(mmol/dL)	1.09±0.30	1.13±0.34	0.27		

\*Data presented as mean ± standard deviation or n (%). HCV = hepatitis C virus; BMI = body mass index; ALT = alanine aminotransferase;AST=aspartate aminotransferase ;FPG = fasting plasma glucose; HDL = high-density lipoprotein.

with A	nti-HCV					
		Crude OR(95%CI)	Adjusted OR(95%CI)	p-value		
Hypertension	Postive=1,Negative=0	0.57(0.29-1.10)	0.62(0.32-1.21)	0.09		
Diabetes Mellitus	Postive=1,Negative=0	0.45(0.19-1.05)	0.52(0.22-1.23)	0.06		
Hyperlipidemia	Postive=1,Negative=0	0.45(0.06-3.33)	0.46(0.06-3.43)	0.65		
Hepatitis	Postive=1,Negative=0	1.54(0.46-5.20)	2.06(0.60-7.11)	0.4		
Drinking	Postive=1,Negative=0	1.73(0.51-5.88)	1.87(0.53-6.58)	0.38		
Smoking	Postive=1,Negative=0	1.56(0.72-3.38)	1.64(0.72-3.76)	0.64		
Betel nut chewing	Postive=1,Negative=0	7.55(2.51-22.74)	9.12(2.96-28.18)	<0.01		
Exercise	Postive=1,Negative=0	1.24(0.76-2.04)	1.01(0.60-1.69)	0.39		
Milk drinking	Postive=1,Negative=0	1.63(1.00-2.65)	1.39(0.84-2.31)	0.05		
Odds ratios were adjusted for age, gender						

 Table 2. Multivariate logistic regression analyses of variables associated