



## The Investigation on the Correlation between Obesity Indicator and Hepatitis B and C

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### Authors' contributions

This work was carried out in collaboration between all authors. All authors read and approved the final manuscript.

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### ABSTRACT

**Background:** Obesity is an important health issue worldwide, and hepatitis B virus (HBV) and hepatitis C virus (HCV) infections are the two major causes of liver disease that lead to Taiwan's medical health and socio-economic problems. There are currently few studies in the nation on the correlation between obesity indicators and hepatitis B and C.

**Purpose:** This study uses adult health check data analysis to understand the correlation of obesity indicators and hepatitis B and C.

**Methods:** This study is a cross-sectional research. The study collected people who did health examinations of a regional hospital in Kaohsiung from 2011 to 2016. The waist circumference (WC), body mass index (BMI), and waist-height ratio (WHR) are used as obesity indicators.

**Results:** A total of 16,459 cases were included in the analysis. The prevalence of abnormal WC is 20.5%, and the WHR abnormal prevalence rate is 32.1%. Underweight Body Mass Index (BMI)  $\text{BMI} \leq 18.4 \text{ kg/m}^2$  (3.8%), normal BMI ranging from 18.5-23.9  $\text{kg/m}^2$  (48.1%), overweight BMI

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ranging from 24.0-26.9 kg/m<sup>2</sup> (26.7%), obesity BMI ≥ 27 kg/m<sup>2</sup> (21.4%). The abnormal rate of hepatitis B was 13.6%, and the abnormal rate of hepatitis C was 1.9%. Logistic regression analysis shows that WC is a risk factor for hepatitis B (OR=1.181, 95%CI=1.014-1.377), and WHR is a protective factor (OR=0.771, 95%CI=0.673-0.885). WHR is a risk factor for hepatitis C (OR=1.571, 95%CI=1.246-1.981).

**Conclusions:** The WC and WHR are respectively the risk factors for hepatitis B and hepatitis C, and the WHR is the protective factor for hepatitis B.

**Keywords:** *Waist circumference (WC); Waist-height ratio (WHR); Body mass index (BMI); hepatitis B; hepatitis C.*

## 1. INTRODUCTION

Taiwan is an area with a high prevalence of hepatitis B, and the carrier rate in general population is about 15% to 20%. The prevalence of hepatitis C is about 4%. Chronic hepatitis B and C are the main causes of liver cancer. According to the survey, about 70% of patients who die of liver cancer are those with hepatitis B and 20% are those with chronic hepatitis C infection. The risk of HCC is increased by more than 100 times if carriers of hepatitis B virus (HBV) or hepatitis C virus (HCV) patients also suffer from obesity and diabetes, there is a multiplication effect. It is pointed out that both obesity and diabetes are risk factors for predicting hepatocellular carcinoma (HCC), and with the differences of infection status of HBV and HCV, it will affect the prevention of HCC [1].

Previous studies on chronic liver disease and obesity have shown in Mexico, the prevalence of chronic liver disease is increasing [2,3] with obesity, diabetes, and metabolic syndrome (MS). Weight gain and diabetes or MS were significantly associated with the increased risk of alanine aminotransferase (ALT) [2]. It is estimated that by 2050, 90% of chronic liver disease cases in Mexico are caused by obesity and alcohol consumption compared with other countries with higher rates of liver disease caused by HBV or HCV [4].

HBV or HCV infection and consuming alcohol are both confirmed risk factors for chronic liver disease [5,6]. Other risk factors include obesity [7,8], MS [9,10] and diabetes [8,11,12]; and the mechanism is developed through nonalcoholic fatty liver disease (NAFLD) and nonalcoholic steatohepatitis (NASH) [8,13,14]. The ratio of chronic liver disease increases rapidly in Mexico with the prevalence of obesity, MS, and diabetes. An article on patients in a teaching hospital in southern Taiwan with chronic hepatitis B (CHB), chronic hepatitis C (CHC) and NAFLD, the

results showed that elevated BMI is an independent risk factor for LC (liver cirrhosis) in three different chronic liver diseases. Therefore, for these patients, weight loss may be beneficial [15]. Other studies have pointed out that obesity, diabetes and hyperlipidemia have recently become potential cofactors for the development of chronic HCV and HBV cases developing into fibrosis [16-18]. At the same time, another Hong Kong study reported that patients with CHB with metabolic syndrome had a higher prevalence of liver cirrhosis than patients with CHB without metabolic syndrome. (38% vs 11%, P <0.001) [19]. Another prospective study from Taiwan, including 2,903 HBsAg-positive men, its median is 14.7 years; higher BMI at baseline is associated with the incidence of NAFLD and liver cirrhosis [20]. A previous study measured visceral obesity by abdominal CT and indicated that HCV infection is a risk factor for the development of insulin resistance, especially in visceral obese patients [21].

In Western countries, 75 to 90% of primary HCC are associated with chronic liver disease [22]. The most common chronic liver disease that causes HCC is hepatitis B or C virus infection and excessive alcohol consumption. Whether the development of HCC is associated with obesity and diabetes or changes in NAFLD is still unclear [23].

There are currently few studies in Taiwan on the correlation between obesity indicator (WC, WHR, BMI) and hepatitis B and C. Therefore, this study uses adult health examination data analysis to understand the correlation between obesity indicator and hepatitis B and C.

## 2. METHODS

### 2.1 Study Design

This study is designed as a cross-sectional study, collecting physical examination and blood

test data as analytical data from people who had a health examination from 2011 to 2016 in a regional hospital in Kaohsiung. All participants were above 20 years of age and met fasting for the examinations.

Inclusion criteria: Those who participated in adult health examination from 2011 to 2016 as subjects.

Exclusion criteria: Age <20 years old and those who had incomplete blood test data and repeated screening are deducted.

## 2.2 Definition of Variables

Height and weight data were obtained using standardised techniques and equipment.

### 1. Definition of obesity indicator:

(1) Waist circumference (WC) outlier: Male  $\geq 90$  cm, female  $\geq 80$  cm. WC was measured at the midpoint between the bottom of the rib cage and the top of the iliac crest.

(2) Waist-height ratio (WHR): Normal ( $< 0.5$ ), abnormal ( $\geq 0.5$ ).

WHR was calculated as WC divided by height.

(3) Body Mass Index (BMI): Taiwan Ministry of Health and Welfare's Standard Classification BMI for 2004.

Underweight: BMI  $\leq 18.4$  kg/m<sup>2</sup>

Normal: BMI between 18.5-23.9 kg/m<sup>2</sup>

Overweight: BMI between 24.0-26.9 kg/m<sup>2</sup>

Obesity: BMI  $\geq 27$  kg/m<sup>2</sup>

### 2. Chronic hepatitis B, C:

After blood biochemical tests, the gastrointestinal specialist judged that it is the asymptomatic carrier of hepatitis B and C.

## 2.3 Ethical Considerations

Data collection of this study began after approval by the hospital's Institutional Review Board (IRB).

## 2.4 Data Processing and Statistical Analysis

All statistical analyses were performed using SPSS software (IBM SPSS Statistics 20; Asia

Analytics Taiwan Ltd., Taipei, Taiwan). Statistical methods include: Descriptive statistics (number of frequencies, percentage, mean and standard deviation), analytical statistics: logistic regression. The above are used to analyse the effects of obesity indicators on hepatitis B and C. Statistically significant level with  $\alpha=0.05$ , and with 95% confidence interval (CI).

## 3. RESULTS

This study includes the analysis from year 2011 to 2016, with 16,459 cases included in the analysis. The result of Table 1 shows that: obesity indicator defines (1) the prevalence of abnormal waist circumference (male:  $\geq 90$  cm, female:  $\geq 80$  cm) 20.5%. (2) Prevalence of abnormal waist-height ratio is 32.1%. (3) Body Mass Index: according to the Health and Welfare Department's standards for Body Mass Index (BMI) in 2004, underweight BMI  $\leq 18.4$  kg/m<sup>2</sup> (3.8%), normal: BMI between 18.5-23.9 kg/m<sup>2</sup> (48.1%), overweight: BMI between 24.0-26.9 kg/m<sup>2</sup> (26.7%), obesity: BMI  $\geq 27$  kg/m<sup>2</sup> (21.4%). The abnormal rate of hepatitis B was 13.6%, and the abnormal rate of hepatitis C was 1.9%.

Logistic regression analysis was performed respectively for the positive or negative of hepatitis B and hepatitis C. The variables included in regression analysis are: gender, age, BMI, waist circumference, waist-height ratio. Table 2 shows that waist circumference is a risk factor for hepatitis B (OR=1.181, 95%CI=1.014-1.377), and waist-height ratio is protective factor (OR=0.771, 95%CI=0.673-0.885). Table 3 shows that waist-height ratio is the risk factor of hepatitis C (OR=1.571, 95%CI=1.246-1.981).

## 4. DISCUSSION

Hepatitis virus infection is a progressive disease that leads to the development of cirrhosis and even hepatocellular carcinoma (HCC); there are about 20  $\pm$  30% of patients worldwide [24,25]. HBV and HCV infection are the two major causes of liver disease that leads to health and socio-economic problems in Taiwan [26,27]. Seventy-five percent of all chronic HBV infections occur in Asia. The prevalence in Taiwan is 15%-20%, and >90% of adults have been infected with hepatitis B virus in the past. It is estimated that there are two million to three million HBV carriers in Taiwan today [28].

**Table 1. Descriptive statistics of demographic characteristics, obesity indicators and Hepatitis B, C (n=16459)**

<b>Variables</b>	<b>Number of people</b>	<b>Percentage</b>	<b>Mean ± standard deviation</b>
<b>Gender</b>			
Male	8987	54.6	
Female	7472	45.4	
<b>Age</b>			
<40 years old	5735	34.8	45.4±11.4
40 years old and above	10724	65.2	
<b>Waist circumference</b>			
Male<90 cm, female<80 cm	13092	79.5	77.7±10.9
Male≥90 cm, female≥80 cm	3367	20.5	
<b>BMI</b>			
< 27kg/m <sup>2</sup>	12940	78.6	24.3±3.9
≥ 27kg/m <sup>2</sup>	3519	21.4	
<b>BMI</b>			
≤ 18.4kg/m <sup>2</sup>	627	3.8	
18.5-23.9kg/m <sup>2</sup>	7918	48.1	
24.0-26.9kg/m <sup>2</sup>	4395	26.7	
≥ 27kg/m <sup>2</sup>	3519	21.4	
<b>Waist-height ratio</b>			
Normal<0.5	11170	67.9	
Abnormal≥0.5	5289	32.1	
<b>Hepatitis B</b>			
Negative	14220	86.4	
Positive	2239	13.6	
<b>Hepatitis C</b>			
Negative	16140	98.1	
Positive	319	1.9	

**Table 2. Regression analysis of obesity indicators on hepatitis B (n=16459)**

<b>Variables<sup>#</sup></b>	<b>β</b>	<b>wald</b>	<b>OR(95%CI)</b>	<b>P value</b>
Gender(female)	0.193	16.363	1.213(1.105-1.332)	<.001
Age(<40 years old)	0.096	3.872	1.101(1.000-1.211)	0.049
WC(normal)	0.167	4.548	1.181(1.014-1.377)	0.033
WHR(normal)	-0.260	13.795	0.771(0.673-0.885)	<.001

Note 1: Stepwise regression method, the variables included in the regression analysis are: gender, age, BMI, WC, WHR.

Note 2: Dependent variable (1) with hepatitis B, (0) without hepatitis B.

<sup>#</sup>( ) is indicated as the reference group

**Table 3. Regression analysis of obesity indicators on hepatitis C (n=16459)**

<b>Variables<sup>#</sup></b>	<b>β</b>	<b>wald</b>	<b>OR(95%CI)</b>	<b>P value</b>
Gender(female)	-0.254	4.765	0.776(0.618-0.974)	0.029
Age(<40 years old)	0.568	17.336	1.766(1.351-2.307)	<.001
WHR(normal)	0.452	14.567	1.571(1.246-1.981)	<.001

Note 1: Stepwise regression method, the variables included in the regression analysis are: gender, age, BMI, WC, WHR.

Note 2: Dependent variable (1) with hepatitis C, (0) without hepatitis C.

<sup>#</sup>( ) is indicated as the reference group

According to data from the Liver Disease Prevention and Treatment Research Foundation, among adults over the age of 20, the prevalence of HCV in Taiwan is estimated at 4.4% (or

423,283 anti-HCV positive carriers) [27]. The study analysed 157,720 patients between 1996 and 2005, the infection rates were similar between males and females, with significant

increases in age and geographic differences. Although the prevalence in most countries is between 1% and 2%, the prevalence in some countries is relatively high, including Egypt (15%), Pakistan (4.7%) and Taiwan (4.4%). The global prevalence of hepatitis C virus (HCV) is about 2% -3%. Between 1990 and 2005, the prevalence of positive anti-HCV antibodies increased from 2.3% to 2.8% [29]. HCV infection causes 60%-80% of those who were infected to develop chronic hepatitis [30] and it is associated with liver steatosis, fibrosis, cirrhosis and hepatocellular carcinoma [31]. The abnormal rate of hepatitis B in the study was 13.6%, and the abnormal rate of hepatitis C was 1.9%, both are lower than the average domestic populace. It may be different because this is a non-national sample survey that it only shows the results of health examination data in a regional hospital.

Previous studies have highlighted the important role of hepatitis virus infection in interacting with obesity. Hepatitis virus infections such as HCV, HBV and HCV/HBV co-infection are positively correlated with the increase in percent body fat (PBF), especially for male [32]. Logistic regression analysis was performed in this study, on whether or not patients have hepatitis B and whether or not they have hepatitis C. The variables included in the regression analysis model are: gender, age, BMI, WC, WHR. It shows that WC is a risk factor for hepatitis B (OR=1.181, 95%CI=1.014-1.377), and WHR is protective factor (OR=0.771, 95%CI=0.673-0.885). The WHR is a risk factor for hepatitis C (OR=1.571, 95% CI=1.246-1.981). Previous studies show that elevated BMI was an independent risk factor associated with possible liver cirrhosis (LC) across the three different etiologies of chronic liver disease. Therefore, weight loss can be beneficial for the patients [15]. Another study points out that WHR may be a better obesity indicator on identifying the individual risk for non-alcoholic fatty liver disease in Korean women [33]. Since previous studies used less of the three obesity indicators: WC, WHR, and BMI respectively on the effects on hepatitis B and C, therefore, it is difficult to compare directly in the literature comparison. However, some studies have shown that obesity is indeed associated with chronic hepatitis B and C and is associated with nonalcoholic fatty liver disease and metabolic diseases. As previous literature has shown, obesity is significantly associated with NAFLD, and visceral fat is more directly related to the onset of NAFLD [34]. Compared with BMI, abdominal obesity is

considered a better predictor of CVD and metabolic diseases. WC has become a widely used measurement method for quantifying abdominal fat accumulation. Epidemiological studies have shown that WHR appears to be more strongly associated with obesity-related diseases and metabolic risk factors than other obesity indicators [35,36]. Since obesity is associated with many diseases and the deterioration of the disease, this study hopes to prevent obesity by finding the correlation between obesity indicators and HBV and HCV, which may help to reduce the progressive deterioration of HBV and HCV. There is also literature [15] pointed out that weight loss can help with the progression of the chronic liver disease. We look forward to the future follow-up study to assess the effectiveness of weight loss to help us understand.

This study had several limitations. First, the study was cross-sectional in design, and hence causal relationships cannot be inferred. Second, this study can only present demographic characteristics, obesity indicators, biochemical blood tests and the correlation between hepatitis B and C. Due to the use of health examination data to perform analysis, the potential impact factors affecting the above results cannot be fully collected, so it is also necessary to be conservative in inference.

## 5. CONCLUSION

HBV and HCV is an important health issue in Taiwan. In particular, hepatitis virus infection is a progressive disease that leads to the development of cirrhosis and even HCC. And liver cancer has been ranked second in the top ten cancers. Obesity is highly associated with many chronic diseases, and is even one of the risk factors for some cancers, such as colorectal cancer, endometrial cancer, and breast cancer. Therefore, if we can find out the correlation between obesity indicators and HBV, HCV, prevention of obesity may help reduce the progressive deterioration of HBV and HCV.

This study shows that waist circumference is a risk factor for hepatitis B, while waist-height ratio is a protective factor. The waist-height ratio is a risk factor for hepatitis C.

## CONSENT

As per international standard or university standard, patient's consent has been collected and preserved by the authors.

## **ETHICAL CONSIDERATIONS**

Data collection of this study began after approval by the hospital's Institutional Review Board (IRB).

## **COMPETING INTERESTS**

Authors have declared that no competing interests exist.

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