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### Discordant Risk: Overweight and cardiometabolic risk in Chinese adults

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#### Abstract

Recent US work identifies "metabolically healthy overweight" and "metabolically at risk normal weight" individuals. Less is known for modernizing countries with recent increased obesity. Fasting blood samples, anthropometry and blood pressure from 8.233 adults aged 18–98 in the 2009 nationwide China Health and Nutrition Survey, were used to determine prevalence of overweight (Asian cut point, BMI 23 kg/m<sup>2</sup>) and five risk factors [pre-diabetes/diabetes (HbA1c 5.7%) inflammation (hsCRP 3 mg/L), pre-hypertension/hypertension (SBP/ DBP 130/85 mmHg), high triglycerides (150 mg/dL), low high-density lipoprotein cholesterol (<40 (men)/<50 mg/dL (women)]. Sex-stratified, logistic and multinomial logistic regression models estimated concurrent obesity and cardiometabolic risk, with and without abdominal obesity, adjusting for age, smoking, alcohol consumption, physical activity, urbanicity, and income. Irrespective of urbanicity, 78.3% of the sample had 1 elevated cardiometabolic risk factor (normal weight: 33.2% had 1 elevated risk factor; overweight: 5.7% had none). At age 18– 30, 47.4% had no elevated risk factors, which dropped to 6% by age 70, largely due to age-related increase in hypertension risk (18–30 years: 11%, >70 years: 73%). Abdominal obesity was highly predictive of metabolic risk, irrespective of overweight (e.g., "metabolically at risk overweight" relative to "metabolically healthy normal weight" [men: Relative Risk Ratio (RRR) = 39.06; 95% Confidence Interval (CI): 23.47, 65.00; women: RRR=22.26; 95% CI: 17.49, 28.33]). To conclude, a large proportion of Chinese adults have metabolic abnormalities. High hypertension risk with age, irrespective of obesity underlies the low prevalence of metabolically healthy

Conflict of Interest

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overweight. Screening for cardiometabolic-related outcomes dependent upon overweight will likely miss a large portion of the Chinese at-risk population.

#### INTRODUCTION

In the US, approximately 16.3 million adults are of normal weight but have two or more metabolic abnormalities.(1) Understanding heterogeneity in risk by overweight status is important particularly in the metabolically at risk but non-overweight population as these individuals often elude screening as they are not perceived as high risk. However, little is known about heterogeneity in risk in Asian populations, and particularly in China where rates of obesity-related non-communicable diseases have recently increased and are now the leading causes of morbidity, disability, and mortality.(2, 3)

Given that Asians are known to have relatively high abdominal obesity(4) and high prevalence of cardiometabolic risk factors at relatively low BMI(5) and at younger ages,(6) understanding how abdominal obesity associates with cardiometabolic risk in the absence of (and in combination with) obesity is important. Although abdominal obesity does not confer the same level of risk for hypertension, diabetes, inflammation and dyslipidemia, many studies rely on syndrome-based indicators (e.g., metabolic syndrome(7, 8), Framingham risk score(9, 10)) and thus may miss important variation that can be observed by investigating variation in co-occurrence of obesity and cardiometabolic risk across individual risk factors.

Understanding the distribution of cardiometabolic risk by age and across overweight and non-overweight individuals is essential for determining the impending burden of cardiovascular disease in China, home to more than 1.3 billion people and comprising one fifth of the world's population. The distribution of cardiometabolic risk factors, particularly among the non-overweight population is particularly important for prevention as these individuals are likely to be missed in screening efforts. To this end, a national Chinese cohort was examined to determine the prevalence of "metabolically healthy overweight" and "metabolically at risk normal weight" by age. Analyses were structured to test the hypothesis that the association between overweight and five risk factors [pre-diabetes/ diabetes (HbA1c 5.7%) inflammation (hsCRP 3 mg/L), pre-hypertension/hypertension (SBP/DBP 130/85 mmHg), high triglycerides ( 150 mg/dL), low high-density lipoprotein cholesterol (<40 (men)/<50 mg/dL (women)] varies, depending upon the indicator. Further analyses tested independent and interactive effects of overweight and abdominal obesity across pre-hypertension or hypertension, pre-diabetes or diabetes, high triglycerides, low HDL-C, and inflammation.

#### MATERIALS & METHODS

#### The China Health and Nutrition Survey (CHNS)

The CHNS collected health data in 228 communities [nine diverse provinces (Guangxi, Guizhou, Heilongjiang, Henan, Hubei, Hunan, Jiangsu, Liaoning, and Shandong)] throughout China from 1989–2009 with eight rounds of surveys, with the 2009 survey collecting fasting blood for the first time. Using multistage, random cluster sampling, counties in the nine provinces were stratified by income and weighted sampling was used to randomly select four counties in each province. Villages and small towns within counties and urban and suburban neighborhoods within cities were selected randomly into primary sampling units that were politically and geographically classified based on State Statistical Office definitions. The surveyed provinces represent 56% of the Chinese population. Survey procedures have been described elsewhere.(11) The study was approved by the Institutional Review Board at the University of North Carolina at Chapel Hill, the China-Japan

#### Analysis Sample

A total of 10,038 adult respondents were surveyed at the 2009 exam, 1,423 did not give blood, 24 were not fasting, and 62 were pregnant, resulting in a total of 8,529 individuals with fasting blood samples. Of these, 237 were missing anthropometry and 60 were missing lab results, resulting in an analytic sample of 8,233 with anthropometry and clinical exam data. There were no statistically significant differences in the total 2009 sample versus the analytic sample in income, urbanization, BMI, cardiometabolic risk or overweight, although the analytic sample included more women, was slightly younger, and had slightly larger waist circumference.

#### Measures

**Cardiometabolic Risk**—Following an overnight fast, blood was collected by venipuncture (12-ml). Whole blood was immediately centrifuged and serum glucose testing completed. The remaining plasma and serum samples were immediately frozen and stored for later laboratory analysis. All samples were processed in a national central lab in Beijing (medical laboratory accreditation certificate ISO 15189:2007) using automatic analyzer, with strict quality control.

Hemoglobin A1c (HbA1c) was measured in whole blood with an automated glycohemoglobin analyzer by using high-performance liquid chromatography system (model HLC-723 G7; Tosoh Corporation, Tokyo, Japan). HbA1c is used as a marker of prediabetes or diabetes (values 5.7%) according to an International Expert Committee.(12) We also conducted comparative sensitivity analyses using fasting glucose (Supplemental Tables 2 and 3, Supplemental Figure 1). Plasma triglyceride and HDL-C were measured by using the glycerol-phosphate oxidase method, the PEG-modified enzyme HDL-C assay by determiner regents (Kyowa Medex Co., Ltd, Tokyo, Japan) on the Hitachi 7600 automated analyzer (Hitachi Inc., Tokyo, Japan). The International Diabetes Federation (IDF) criteria for adults(13) is used for indicators of dyslipidemia risk (high triglycerides: 150 mg/dL; low HDL-C: (male: <40 mg/dL; female<50 mg/dL) or taking lipid lowering medication. High-sensitivity C-reactive protein (hsCRP), using the immunoturbidimetric method with Denka Seiken, Japan reagents (Hitachi 7600 automated analyzer, Hitachi Inc., Tokyo, Japan) is used as a marker of inflammation, with values 3 mg/dL indicating high risk.(14) Elevated hsCRP is a strong independent risk factor for future cardiovascular events, regardless of the underlying condition.(15)

Systolic (SBP) and diastolic (DBP) blood pressure were measured on the right arm, using mercury sphygmomanometers with appropriate cuff sizes. Measures were collected in triplicate after a 10 minute seated rest and the mean of the three measurements used in analyses. Risk was determined as systolic blood pressure (SBP) 130mmHg/diastolic blood pressure (DBP) 85mmHg according to the IDF cut point for pre-hypertension or taking blood pressure medication.(16)

**Overweight and Abdominal Obesity**—Height was measured without shoes to the nearest 0.2 cm using a portable SECA stadiometer, and weight was measured without shoes and in light clothing to the nearest 0.1 kg on a calibrated beam scale. Overweight was defined using the recommended Asian BMI cut point of 23 kg/m<sup>2</sup>, with non-overweight <23 kg/m<sup>2</sup>.(17) We also conducted comparative sensitivity analyses using the WHO BMI 25 kg/m<sup>2</sup> cut point(18) (Supplemental Tables 4 and 5, Figure 2). Abdominal obesity

was classified using waist circumference, which was measured with a SECA tape measure midway between the between the lower rib margin and the iliac crest. The IDF recommends sex and ethnic-specific waist circumference cut points, thus the Asian cut point for abdominal obesity for adults (waist circumference: men 90 cm; women 80 cm) was used. (13) Abdominal obesity is associated with cardiometabolic risk, independent of BMI,(19) particularly in individuals with normal BMI.(20, 21)

**Metabolically Healthy and Overweight Categories**—Cardiometabolic risk is defined as having elevated values of one or more of the following five risk factors: inflammation (hsCRP 3 mg/L), pre-hypertension or hypertension (SBP/DBP 130/85 mmHg), high triglycerides (triglycerides: 150 mg/dL), low HDL-C: (male: <40mg/dL; female<50 mg/ dL), and pre-diabetes or diabetes (HbA1c 5.7%). Patterns of risk associated with overweight were used to define the following four categories: "metabolically healthy, nonoverweight" (BMI< 23 kg/m<sup>2</sup>, no risk factors); "metabolically healthy, overweight" (BMI 23 kg/m<sup>2</sup>, no risk factors); "metabolically at risk, non-overweight" (BMI< 23 kg/m<sup>2</sup>, 1 risk factor); and "metabolically at risk, overweight" (BMI 23 kg/m<sup>2</sup>, 1 risk factor); and "metabolically at risk, overweight" (BMI 23 kg/m<sup>2</sup>, 1 risk factor). Inflammation was considered part of the cardiometabolic risk profile since in developing countries, high rates of obesity occur alongside high rates of infectious disease and malnutrition, resulting in dual burden of obesity-related and pathogen-related inflammation(22) and future cardiovascular disease.(23) Abdominal obesity was not considered in the definitions of these phenotypes due to its covariation with BMI and conceptual interest in the association of abdominal obesity with these phenotypes.

**Control variables**—Age was recorded as the respondent's age on the date of exam, ranging from 18 to 98 years and categorized into the following categories: 18–30, 30–40, 40-50, 50-60, 60-70, and >70 years. Number of cigarettes per day were reported and categorized as currently smoking one or more cigarettes per day. Alcohol was reported as number of alcoholic beverages consumed per week across a variety of beverage types, categorized according to consumption of 1 servings of alcohol per day. Physical activity was reported using a detailed 7-day physical activity recall instrument and leisure activities assigned Metabolic Equivalent (MET) values using the Compendium of Physical activity, (24) which has been used in international contexts, including China.(25) MET values were multiplied by the time spent in each activity and categorized according to 27 METs per week, equivalent to 1 hour per week of moderate-intensity, such as brisk walking. Urbanicity was defined using a multidimensional 12 component urbanization index capturing community-level physical, social, cultural, and economic environments and represents the heterogeneity otherwise missed in an urban/rural measure based only on population density.(26) Tertiles of the 100 value index represent low, medium, and high urbanicity. Household assets and household income were used to derive income; tertiles represent low, medium, and high income.(27)

#### **Statistical Analyses**

We conducted statistical analyses using Stata (Release 11.0, Stata Corporation, College Station, TX). In descriptive analyses, markers are compared across overweight status (percentages for categorical variables, means for continuous variables). A series of models were used to examine overweight (BMI 23 kg/m2) and cardiometabolic risk patterning (1 of the following risk factors: inflammation, pre-hypertension or hypertension, high triglycerides, low HDL-C, pre-diabetes or diabetes) and assesses the independent and interactive effects of overweight and abdominal obesity (waist circumference: men 90 cm; women 80 cm). First, multinomial regression models predicting risk of falling into one of four categories: 1) "metabolically healthy, normal weight", 2) "metabolically healthy, overweight", 3) "metabolically at risk, overweight," and 4) "metabolically at risk, normal

weight"[referent], with abdominal obesity as the main exposure were used to test the hypothesis that abdominal obesity would differentially predict membership into one of the four risk categories. Second, a series of five separate logistic regression models for each outcome (inflammation, pre-hypertension or hypertension, high triglycerides, low HDL-C, pre-diabetes or diabetes) were used sequentially to test the hypotheses that: 1) overweight (first main exposure) differentially associates with each cardiometabolic risk factor, and 2) abdominal obesity (second main exposure), controlling for overweight, confers differential risk for each cardiometabolic marker. Third, five separate and sequential logistic regression models for each outcome (inflammation, pre-hypertension or hypertension, high triglycerides, low HDL-C, pre-diabetes or diabetes) with four main exposure variables: 1) overweight with abdominal obesity; 2) overweight with no abdominal obesity; 3) nonoverweight with abdominal obesity; and 4) non-overweight with no abdominal obesity [referent], were used to test the hypothesis that there is an interactive effect between overweight and abdominal obesity, with overweight and abdominal obese individuals at highest risk. All models were sex-stratified and included the following covariates: age, smoking, alcohol consumption, physical activity, urbanicity, and income. While model building suggested minimal change in effect of the main exposure with the addition of smoking, alcohol consumption, physical activity, urbanicity, and income, these covariates were retained on conceptual grounds. Two forms of sensitivity analyses were performed: (1) using the BMI 25 kg/m<sup>2</sup> cut point rather than the more stringent Asian BMI 23 kg/m<sup>2</sup> cut point, and (2) using impaired fasting glucose ( 100 mg/dL) as opposed to HbA1c (prediabetes or diabetes) as the indicator of diabetes risk. Results from both sensitivity analyses are shown in the Supplemental Materials.

#### RESULTS

While approximately half of the sample was overweight, 78% had elevated values of one or more cardiometabolic risk factors (49% had more than two) (Table 1). The proportion with pre-hypertension or hypertension (42.8%) and pre-diabetes or diabetes (38.1%) was particularly high. Rates of isolated hypertension risk (9.9%) and pre-diabetes or diabetes (7.7%) were roughly twice as high as isolated high triglycerides, low HDL-C, and inflammation, which were around 4%. In addition, mean waist circumference of the metabolically healthy non-overweight group were on average 16 cm less than that of the metabolically at risk overweight group, 10 cm less than the metabolically healthy overweight group.

All cardiometabolic risk factors were higher among the overweight (Table 2). The proportion of overweight and non-overweight individuals with inflammation, prehypertension or hypertension, and pre-diabetes or diabetes increased with age, whereas rates of both high triglycerides and low HDL-C were highest in the early adult years for men and HDL-C was relatively stable among women. Relative to the other risk factors, differences in hypertension risk by overweight status were not as large. Mean continuous values for each marker are shown in Supplemental Table 1.

A comparatively large proportion of the sample across all age groups was "metabolically at risk normal weight", notably 42% of the total sample aged 18–30 (Figure 1). There were very few "metabolically healthy overweight" individuals and the proportion of the sample considered metabolically healthy (regardless of overweight status) declined considerably with age. Findings were similar across low, medium, and high urbanicity.

We used multinomial regression models to test the hypothesis that abdominal obesity differentially predicted membership into one of the four categories of obesity with and without metabolic risk (Table 3). The odds of being in the "metabolically at risk

overweight" compared to the metabolically healthy, non-overweight group was dramatically elevated with abdominal obesity (men, Odds Ratio (OR): 39.06, 95% Confidence Interval (CI): 23.47–65.00; women, OR: 22.26, CI: 17.49–28.33). Abdominal obesity was also predictive of having high metabolic risk among both non-overweight and overweight individuals.

We used cardiometabolic risk factor-specific logistic regression models to examine the association between overweight and each risk factor, with and without control for abdominal obesity (Set A, models 1 and 2). One set of models included overweight exposure only (Table 4, Column 2), a second set of models included both overweight and abdominal obesity as exposure variables (Table 4, Column 3). As expected, overweight independently increased the risk of each cardiometabolic marker in men and women, with variation across indicators. Independent of overweight status, abdominal obesity conferred risk for all cardiometabolic indicators.

A third set of cardiometabolic risk factor-specific logistic regression models was used to examine the interactive association between overweight and abdominal obesity with each risk factor (Set B models): 1) overweight with abdominal obesity, 2) overweight with no abdominal obesity, and 3) non-overweight with abdominal obesity relative to 4) non-overweight with no abdominal obesity. Overweight individuals with abdominal obesity were at comparatively high cardiometabolic risk, with variation across pre-hypertension or hypertension, pre-diabetes or diabetes, inflammation and dyslipidemia (Table 4, Column 6). In the majority of cases, there was close to or above a threefold higher risk with the combination of overweight and abdominal obesity relative to non-overweight and no abdominal obesity.

#### **Sensitivity Analysis Results**

Results were remarkably similar when impaired fasting glucose ( 100 mg/dL) as opposed to HbA1c (pre-diabetes or diabetes) was used as the indicator of diabetes risk (Supplemental Tables 2 and 3, Supplemental Figure 1). Further, results were similar when overweight was defined using BMI 25 kg/m<sup>2</sup> cut point(18) (Supplemental Tables 4 and 5), but with a considerably larger portion of the population considered "metabolically at risk non-overweight" (Supplemental Figure 2). Relative to the BMI 23 kg/m<sup>2</sup> cut point, using the BMI 25 kg/m<sup>2</sup> cut point suggests that abdominal obesity has the same (albeit stronger) positive association with the metabolically at risk overweight.

#### DISCUSSION

Over three-quarters of this national Chinese sample had elevated levels of at least one cardiometabolic risk factor. Even in young adults, only half were "metabolically healthy" at 18–30 years of age, but this decreased to 6% in the 60–70 year olds. Findings were similar across low, medium, and high urbanicity, indicating high prevalence of metabolic abnormalities even in the rural Chinese population, according to the IDF criteria for metabolic risk. Just over 5% of the overweight adults had no risk factors. Among the non-overweight adults, 33.2% had at least one risk factor. Rates of isolated high triglycerides, low HDL-C, and inflammation were relatively low. The predominant risk factor was hypertension risk, which increased dramatically with age, underlying the decline with age in the proportion of adults considered metabolically healthy. While we cannot rule out the fact that the high rates of hypertension risk might be a result of the normal increase in blood pressure with age, the high prevalence of metabolic abnormalities among the 18–30 year old group, particularly in low HDL-C and high triglycerides, is notable. Our findings highlight the importance of screening in the general population, given the large portion of non-

overweight individuals with cardiometabolic risk. This should be at the forefront in China's efforts to prevent and control non-communicable diseases.

Previous research suggests that hypertension risk is particularly high in China and is found at low BMI levels in Asians relative to other subpopulations. (5, 28, 29) A recent paper on diabetes in China documented high diabetes prevalence, (30) although that sample does not represent poor, less urbanized areas, as well as the CHNS. While HDL-C tends to be relatively low in Asians,(31) rates of low HDL-C were not extraordinary in these Chinese adults. Inflammation, a risk factor for type 2 diabetes and cardiovascular disease,(32-34) is associated with insulin resistance and cardiometabolic risk in the absence of obesity.(35) The observed findings suggest that inflammation increases with age across all adults, which is common in developing countries where high rates of obesity occur alongside high rates of infectious disease and malnutrition, resulting in dual burden of obesity-related and pathogen-related inflammation.(22) Elevated hsCRP is a strong independent risk factor for future cardiovascular events, regardless of the underlying condition.(15) For example, risk may be present even if the underlying etiology relates to periodontal disease, which is an independent risk factor for coronary heart disease.(36) Despite the sex difference in abdominal obesity (largely due to the lower threshold for abdominal obesity in women (80 cm) versus men (90 cm), inflammation rates were similar. The combined association between overweight and abdominal obesity with cardiometabolic risk factors suggests a multiplicative effect on cardiometabolic risk, with risk that is magnified in overweight individuals, even after controlling for smoking, alcohol consumption, total physical activity, urbanicity, and income.

Most cardiometabolic screening is aimed at individuals above a given threshold of body mass, despite the fact that elevated cardiometabolic risk is common in the general population.(37) The observed high risk among non-overweight Chinese, and the increase in risk with age, implies that screening for cardiometabolic risk should occur across all adults, regardless of weight status. Further, the high prevalence of pre-hypertension and hypertension, pre-diabetes or diabetes, high triglycerides, low HDL-C, and inflammation risk even below the BMI 23 kg/m<sup>2</sup> threshold supports a lower BMI cut point for the Chinese population.(4, 17)

China has had a history of undernutrition followed by the most rapid increase in obesity and related diseases worldwide, with differential rates across rural and urban areas.(2, 38) In the last decade, changes in diet and activity in China have occurred more rapidly than previously recorded for any country,(2) including increased intake of edible oil, fried foods, animal source foods and snacking, as well as declining occupational, domestic, and travel activity expenditure and increasing TV time.(25, 39) The observed findings fit with the trend towards less healthier lifestyles, and indicate elevated cardiometabolic risk even in the rural areas of China.

There are a few limitations to this analysis. This cross-sectional analysis does not examine trends in risk or determinants of cardiometabolic risk. The main objective of this study was to determine the prevalence of "metabolically healthy overweight" and "metabolically at risk non-overweight" in Chinese adults and to examine patterns of risk by age and gender. As such, the analytic strategy was designed to adjust for key covariates, rather than undertaking causal modeling of these relationships. The use of conventional, anthropometric and cardiometabolic cut points do not capture differences in cardiometabolic risk that exist on the continuum of BMI and waist circumference. Likewise, cut points do not capture the complex process of cardiometabolic risk However, cut points are needed for clinical guidance and there is rationale for use of Asian-specific BMI cut points based on higher risk at lower BMI.(4, 17, 40) Finally, BMI does not discriminate between lean and adipose tissue

and waist circumference is an indicator of both the subcutaneous and visceral adipose tissue compartments,(41) thus we cannot distinguish the two.

In summary, findings suggest a high prevalence of cardiometabolic risk in China that increases with age. The majority of the sample is considered metabolically at risk, with similar patterns of risk across urbanicity. Rates of risk increase dramatically with age and with overweight and abdominal obesity. The combined association between overweight and abdominal obesity with cardiometabolic risk factors, even after controlling for smoking, alcohol consumption, total physical activity, urbanicity, and income, suggest extraordinary burden of future cardiovascular disease with population increases in weight and abdominal obesity – a major clinical concern. Screening for cardiometabolic-related outcomes dependent upon overweight will likely miss a large portion of the Chinese at-risk population and has implications for other Asian populations.

#### **Supplementary Material**

Refer to Web version on PubMed Central for supplementary material.

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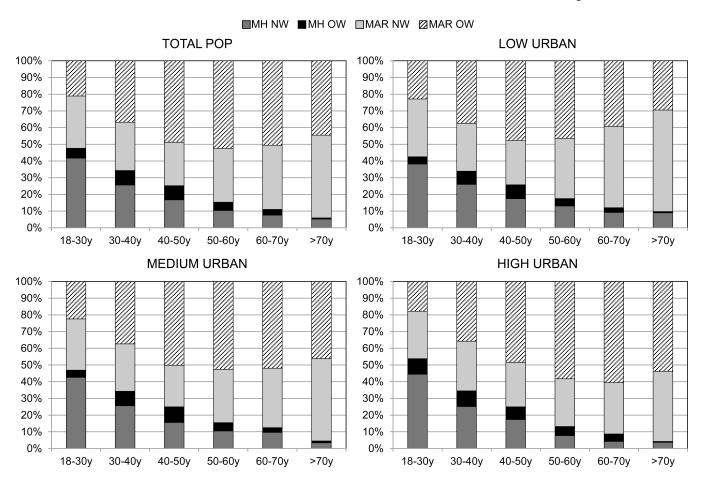
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### Figure 1. Concurrent Overweighta and Cardiometabolic Riskb, Total Population and Across Urbanicity

<sup>a</sup>Overweight defined using the BMI 23 kg/m<sup>2</sup> cutpoint(17)

<sup>b</sup>Cardiometabolic Risk, defined as one or more of the following risk factors: dysglycemia (HbA1c 5.7%) inflammation (**hsCRP** 3 mg/L), **pre-hypertension/hypertension** (SBP/DBP 130/85 mmHg), high triglycerides (150 mg/dL), low high-density lipoprotein cholesterol (<40 mg/dL, men; <50 mg/dL, women).

NOTE: MH NW = Metabolically Healthy, Normal Weight

MH OW = Metabolically Healthy, Overweight

MAR NW = Metabolically at Risk, Normal Weight

MAR OW = Metabolically at Risk, Overweight

## Table 1

Sample Descriptive Statistics, 2009 China Health and Nutrition Survey, mean (SD), percent (95% Confidence Interval)

	Men n=3,866	Women n=4,367	Total n=8,233
Demographics Age (years)	50.8 (15.1)	51.0 (14.9)	50.9 (15.0)
Anthropometrics Body Mass Index (kg/m <sup>2</sup> )	23.3 (3.4)	23.4 (3.5)	23.4 (3.5)
Percent Overweight (BMI 23 kg/m <sup>2</sup> )(17)	51.2 (49.6, 52.8)	50.5 (49.0, 52.0)	50.8 (49.8, 51.9)
Percent Abdominal Obesity (waist circumference: men 90 cm; women 80 cm) (13)	30.4 (29.7, 31.2)	54.6 (53.8, 55.3)	43.2 (42.7, 43.8)
Waist circumference (cm)	84.4 (10.2)	81.2 (10.2)	82.7 (10.4)
Cardiometabolic Risk Percent with Inflammation (hsCRP $3 \text{ mg/L}$ )	23.8 (23.1, 24.5)	23.9 (23.3, 24.6)	23.9 (23.4, 24.3)
Percent with Pre-hypertension or hypertension ( $130/85 \text{ mmHg})^{a}$	46.7 (45.9, 47.5)	39.2 (38.5, 40.0)	42.8 (42.2, 43.3)
Percent with High Triglycerides ( 150 mg/dL) <sup><math>a</math></sup>	35.3 (34.5, 36.1)	29.2 (28.5, 29.8)	32.0 (31.5, 32.6)
Percent with Low HDL-C (male: <40 mg/dL; female<50 mg/dL)	17.8 (17.2, 18.5)	33.7 (33.0, 34.4)	26.3 (25.8, 26.8)
Percent with Pre-diabetes or diabetes (HbA1c 5.7%)	38.4 (37.7, 39.2)	37.7 (37.0, 38.5)	38.1 (37.5, 38.6)
Percent with any of the five risk factors $^{b}$	80.1 (79.5, 80.8)	76.7 (76.0, 77.3)	78.3 (77.8, 78.7)
Percent with two or more of the five risk factors	48.2 (47.7, 50.8)	47.9 (46.4, 49.4)	48.5 (47.5, 49.6)
Percent with Isolated <sup>c</sup> Inflammation (hsCRP 3 mg/L)	4.4 (4.1, 4.7)	3.4 (3.1, 3.7)	3.9 (3.7, 4.1)
Percent with Isolated <sup>c</sup> Pre-hypertension or hypertension ( $130/85 \text{ mmHg})^d$	11.9 (11.4, 12.4)	8.1 (7.7, 8.5)	9.9 (9.5, 10.2)
Percent with Isolated $^b$ High Triglycerides ( $150 \text{ mg/dL})^d$	5.8 (5.4, 6.1)	2.7 (2.5, 3.0)	4.2 (3.9, 4.4)
Percent with Isolated <sup>c</sup> Low HDL-C (male: <40 mg/dL; female<50 mg/dL)	1.6 (1.4, 1.8)	6.6 (6.2, 6.9)	4.2 (4.0, 4.4)
Percent with Isolated <sup>C</sup> Pre-diabetes or diabetes (HbA1c 5.7%)	7.3 (6.9, 7.7)	8.0 (7.6, 8.4)	7.7 (7.4, 7.9)
Weight Status and Cardiometabolic Risk Profiles Metabolically healthy, non-overweight $d$	14.6 (14.0, 15.2)	>17.2 (16.7, 17.8)	>16.0 (15.6, 16.4)
Metabolically healthy, overweight $^{d}$	5.3 (4.9, 5.6)	6.1 (5.7, 6.4)	5.7 (5.4, 6.0)
Metabolically at risk, non-overweight <sup>d</sup> Metabolically at risk, overweight <sup>d</sup>	34.2 (33.4, 35.0) 45.9 (45.1, 46.7)	32.2 (31.5, 32.9) 44.4 (43.7, 45.2)	33.2 (32.6, 33.7) 45.1 (44.6, 45.7)
Waist Circumference across Weight and Cardiometabolic Risk Profiles Waist circumference (cm) in the Metabolically healthy, non-overweight $d$	5.7 (75.4, 76.0)	72.5 (72.2, 72.7)	73.9 (73.7, 74.1)

	Men n=3,866	Women n=4,367	Total n=8,233
Waist circumference $(cm)$ in the Metabolically healthy, overweight <sup>d</sup>	86.7 (86.2, 87.3)	82.2 (81.8, 82.6)	84.2 (83.9, 84.5)
Waist circumference $(cm)$ in the Metabolically at risk, non-overweight $d$	78.4 (78.2, 78.7)	76.1 (75.9, 76.3)	77.2 (77.1, 77.4)
Waist circumference $(cm)$ in the Metabolically at risk, overweight $d$	91.2 (91.1, 91.4)	88.2 (88.0, 88.4)	89.7 (89.5, 89.8)
Waist circumfernce (cm) in those with any of the five risk factors	75.7 (75.4, 76.0)	72.5 (72.2, 72.7)	73.9 (73.7, 74.1)
Behavioral and Social Factors			
Current Smoker	54.8 (0.8)	3.6 (0.3)	27.6 (0.5)
1 Alcohol Serving per day	26.9 (0.7)	1.9 (0.2)	13.6 (0.4)
Highly Active (Leisure Metabolic Equivalents (METs) 27 METs/week) <sup>e</sup>	34.0 (0.8)	26.6 (0.7)	30.1 (0.5)
Urbanicity (100 point modernization index)(24)	67.2 (19.5)	67.7 (19.4)	67.4 (19.5)
Household Income (yuan)	39,086.2 (47,953.0)	37,139.1 (43,028.4) 38,055.7 (45,420.8)	38,055.7 (45,420.8)
a or medication			

 $b_{\rm thermal}$  inflammation, pre-hypertension or hypertension, high triglycerides, low HDL-C, pre-diabetes or diabetes

 $c_{\rm r}$ 

d Metabolically healthy indicates no inflammation, Pre-hypertension or hypertension, high triglycerides, low HDL-C, pre-diabetes or diabetes, metabolically at risk indicates at least one of the five risk factors. Overweight is defined using the recommended Asian BMI curpoint of  $23 \text{ kg/m}^2$ , with non-overweight <23 kg/m<sup>2</sup>.(17)

 ${}^{\!\!\!\mathcal{C}}_{\!\!\!}$  Equivalent to 1 hour of brisk walking per day

Pregnant women excluded

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# Table 2

Sample Descriptive Statistics By Overweight (OW)<sup>a</sup> status (%, 95<sup>th</sup> CI)

Men	Inflammation <sup>b</sup>		Pre-hypertension/ hypertension <sup>b</sup>	/	High Triglycerides $^{b}$	esb	Low HDL-C <sup>b</sup>		Pre-diabetes/diabetes <sup>b</sup>	etes <i>b</i>
	Non-OW N=2,754	OW N=1,154	Non-OW N=2,754	OW N=1,154	Non-OW N=2,754	OW N=1,154	Non-OW N=2,754	OW N=1,154	Non-OW N=2,754	OW N=1,154
18-30y	9.5 (7.6, 11.3)	25.6 (21.7, 29.4)	10.3 (8.3, 12.2)	33.3 (29.2, 37.5)	21.8 (19.2, 24.5)	41.9 (37.5, 46.2)	14.4 (2.3, 12.1)	30.2 (4.1, 26.2)	9.9 (8.0, 11.8)	23.3 (19.5, 27.0)
30-40y	16.5 (14.3, 18.7)	20.5 (18.2, 22.8)	22.5 (20.0, 24.9)	34.7 (32.0, 37.4)	20.0 (17.6, 22.4)	54.6 (51.8, 57.4)	12.3 (1.9, 10.3)	29.7 (2.6, 27.1)	17.9 (15.6, 20.2)	34.4 (31.7, 37.1)
40–50y	15.8 (13.9, 17.8)	25.0 (23.1, 27.0)	26.4 (24.0, 28.8)	51.9 (49.6, 54.1)	24.6 (22.3, 27.0)	56.2 (54.0, 58.4)	10.0 (1.6, 8.3)	23.9 (1.9, 22.0)	28.2 (25.7, 30.6)	40.5 (38.3, 42.7)
50-60y	19.3 (17.5, 21.2)	27.2 (25.2, 29.2)	40.7 (38.3, 43.0)	62.0 (59.9, 64.2)	22.7 (20.7, 24.6)	47.2 (45.0, 49.4)	11.1 (1.5, 9.6)	24.7 (1.9, 22.7)	32.0 (29.8, 34.2)	51.3 (49.1, 53.5)
60-70y	24.0 (21.7, 26.4)	27.4 (24.9, 29.9)	49.5 (46.8, 52.3)	74.8 (72.4, 77.2)	18.5 (16.4, 20.7)	43.4 (40.6, 46.1)	7.9 (1.5, 6.4)	21.5 (2.3, 19.3)	39.5 (36.8, 42.2)	59.1 (56.3, 61.8)
>70y	37.2 (34.1, 40.4)	47.3 (43.6, 51.0)	66.9 (63.9, 70.0)	77.4 (74.3, 80.5)	17.2 (14.7, 19.6)	38.2 (34.6, 41.7)	6.3 (1.6, 4.7)	23.7 (3.1, 20.5)	49.4 (46.1, 52.6)	66.1 (62.6, 69.6)
Women	Inflammation <sup>b</sup>		Pre-hypertension/ hypertension <sup>b</sup>	'n	High Triglycerides $^{b}$	esp	Low HDL-C <sup>b</sup>		${ m Pre-diabetes/diabetes}^b$	oetesb
	Non-OW N=2,754	0W N=1,154	Non-OW N=2,754	OW N=1,154	Non-OW N=2,754	OW N=1,154	Non-OW N=2,754	OW N=1,154	Non-OW N=2,754	0W N=1,154
18-30y	8.9 (7.3, 10.6)	31.2 (25.9, 36.5)	1.3 (0.6, 1.9)	13.0 (9.1, 16.8)	9.6 (7.9, 11.3)	19.5 (14.9, 24.0)	23.6 (2.4, 21.2)	40.3 (5.6, 34.6)	11.8 (10.0, 13.6)	26.0 (20.9, 31.0)
30-40y	9.0 (7.6, 10.5)	22.4 (19.8, 25.1)	7.0 (5.7, 8.3)	21.3 (18.7, 23.8)	8.5 (7.1, 9.9)	29.9 (27.0, 32.8)	25.8 (2.2, 23.6)	45.7 (3.1, 42.5)	15.3 (13.5, 17.1)	25.6 (22.8, 28.3)
40–50y	12.5 (10.9, 14.0)	21.1 (19.4, 22.8)	18.7 (16.9, 20.5)	36.9 (34.9, 38.9)	17.2 (15.5, 19.0)	32.9 (30.9, 34.8)	24.5 (2.0, 22.5)	41.8 (2.1, 39.8)	20.6 (18.8, 22.5)	37.8 (35.8, 39.8)
50-60y	17.1 (15.3, 19.0)	31.5 (29.7, 33.3)	32.6 (30.3, 34.9)	55.1 (53.2, 57.0)	25.5 (23.3, 27.6)	45.3 (43.4, 47.3)	24.0 (2.1, 22.0)	41.6 (1.9, 39.7)	31.0 (28.7, 33.2)	53.3 (51.4, 55.2)
60-70y	27.1 (24.5, 29.7)	41.4 (38.9, 43.9)	52.4 (49.5, 55.4)	70.6 (68.3, 72.9)	27.4 (24.8, 30.1)	47.9 (45.4, 50.4)	28.5 (2.7, 25.8)	45.1 (2.5, 42.6)	43.1 (40.1, 46.0)	61.8 (59.4, 64.3)
>70y	28.6 (25.9, 31.3)	48.5 (45.3, 51.8)	68.1 (65.3, 70.9)	81.6 (79.1, 84.1)	23.6 (21.0, 26.1)	44.4 (41.1, 47.6)	25.4 (2.6, 22.7)	36.0 (3.1, 32.9)	48.2 (45.2, 51.2)	68.6 (65.6, 71.6)
<sup>a</sup> 2009 Chi	na Health and Nutrit	$^{\it a}_{\it 2009}$ China Health and Nutrition Survey, Overweight defined		using the BMI 23 kg/m <sup>2</sup> cutpoint(16)	utpoint(16)					
b pre-diabe cholestero	tes or diabetes (Hb/ l (<40 mg/dL, men;	b pre-diabetes or diabetes (HbA1c 5.7%) inflammation (hsCRI cholesterol (<40 mg/dL, men; <50 mg/dL, women)	tion (hsCRP	3 mg/L), Pre-hypertension or hypertension (SBP/DBP 130/85 mmHg), high triglycerides ( 150 mg/dL), low high-density lipoprotein	n or hypertension (S	BP/DBP 130/85 m	mHg), high triglyce	erides ( 150 mg/dL	.), low high-density l	lipoprotein

## Table 3

Concordant Cardiometabolic and Overweight Relative Risk With Abdominal Obesity<sup>a</sup>

					TE I	
RRR		Z	Ь	RRR	Z	Ъ
Metabolically healthy $^{b}$ non-overweight $^{c}$ 1.0 Abdominal obesity	ı	1	1	1.0	ł	1
Metabolically healthy $^b$ overweight $^c$ Abdominal obesity $^c$ 16.45 (9.21, 29.37)		9.467	<.001	<.001 7.94 (5.73, 11.01)	12.435	<.001
Metabolically at Risk $^b$ non-overweight $^{\cal C}$ 2.25 (1.30, 3.89) Abdominal obesity $^{\cal C}$		2.906	<.01	<.01 1.59 (1.25, 2.01)	3.777	<.001
Metabolically at Risk <sup>b</sup> overweight <sup>c</sup> Abdominal obesity <sup>c</sup> $39.06(23.47, 6)$	55.00) 1	4.105	<.001	39.06 (23.47, 65.00) 14.105 <.001 22.26 (17.49, 28.33) 25.225		<.001
$^{a}$ Relative risk of falling into each category of cardiometabolic risk and overwe weight as the referent category, 2009 China Health and Nutrition Survey. Mode (across leisure, commuting, and occupational activity), urbanicity, and income	lic risk and rition Surv nicity, and	l overwei 'ey. Modε l income	ight for a	bdominal obesity, sex-s for age, current smokir	stratified n ng, drinkin	<sup>a</sup> Relative risk of falling into each category of cardiometabolic risk and overweight for abdominal obesity, sex-stratified multinomial logistic regression model results, with metabolically healthy normal weight as the referent category, 2009 China Health and Nutrition Survey. Models adjust for age, current smoking, drinking 7+ alcoholic beverages per week, engaging in high levels of total physical activity (across leisure, commuting, and occupational activity), urbanicity, and income

3 mg/L), pre-hypertension or hypertension (SBP/DBP 130/85 mmHg), high triglycerides ( 150 mg/dL), low high-density lipoprotein cholesterol (<40 mg/dL, men; <50 mg/dL, women). Metabolically at risk defined as one or more of the above risk factors

 $^{\rm C}$  Overweight is defined using the recommended Asian BMI cutpoint of 23 kg/m<sup>2</sup>, with non-overweight <23 kg/m<sup>2</sup>.(17); Abdominal obesity (waist circumference: men 90 cm; women 80 cm) (13)

NOTE: Model interpretation: Abdominal obesity (versus no abdominal obesity) predicts membership in "metabolically healthy overweight" category at RRR=16.45 (CI 9.21-29.37) relative to membership in the "metabolically healthy non-overweight category"

# Table 4

Odds ratios for overweight: independently, controlling for, or interacted with, abdominal obesity<sup>a</sup>

			Men	u	
Risk factor	Set A Models: 1 (o controlling for abdo	Set A Models: 1 (overweight only <sup>c</sup> ) and 2 (overweight <sup>c</sup> controlling for abdominal obesity <sup>c</sup> ) Odds Ratio (95% $CD^c$	l 2 (overweight <sup>c</sup> s Ratio (95% CI) <sup>c</sup>	Set B Models. Odds Ratio (95% CI) Overweight <sup>al</sup> Interacted with Abdominal Obesity <sup>c</sup>	5% CI) ninal Obesity <sup>c</sup>
		(Set A 1)	(Set A 2)		(Set B)
Inflammation <i>b</i>	Overweight	1.55 (1.32, 1.82)	1.19 (0.99, 1.44)	Non-overweight/No Abdominal Obesity	1.00 (1.00, 1.00)
	Abdominal Obesity	1	1.70 (1.41, 2.05)	Overweight/No Abdominal Obesity	1.19 (0.97, 1.46)
	1	1	;	Non-overweight/ Abdominal Obesity	1.69 (1.08, 2.64)
	1	1	;	Overweight/Abdominal Obesity	2.03 (1.69, 2.44)
Pre-/hypertension b	Overweight	2.56 (2.22, 2.96)	1.99 (1.68, 2.34)	Non-overweight/No Abdominal Obesity	1.00 (1.00, 1.00)
	Abdominal Obesity	1	1.73 (1.45, 2.06)	Overweight/No Abdominal Obesity	1.88 (1.57, 2.24)
	1	I	;	Non-overweight/Abdominal Obesity	1.23 (0.80, 1.88)
	-	-	1	Overweight/Abdominal Obesity	3.48 (2.93, 4.15)
High Triglycerides $b$	Overweight	3.27 (2.83, 3.79)	2.63 (2.23, 3.11)	Non-overweight/No Abdominal Obesity	1.00 (1.00, 1.00)
	Abdominal Obesity	I	1.60 (1.35, 1.89)	Overweight/No Abdominal Obesity	2.65 (2.21, 3.17)
	1	ł	;	Non-overweight/Abdominal Obesity	1.66 (1.07, 2.59)
	-	-		Overweight/Abdominal Obesity	4.21 (3.54, 5.00)
Low HDL-C <sup>b</sup>	Overweight	2.82 (2.34, 3.40)	2.02 (1.63, 2.50)	Non-overweight/No Abdominal Obesity	1.00 (1.00, 1.00)
	Abdominal Obesity	-	1.97 (1.61, 2.42)	Overweight/No Abdominal Obesity	2.05 (1.62, 2.59)
	-	-	-	Non-overweight/Abdominal Obesity	3.99 (3.23, 4.94)
	-	-	-	Overweight/Abdominal Obesity	2.14 (1.26, 3.64)
$\operatorname{Pre-/diabetes} b$	Overweight	2.22 (1.93, 2.56)	1.70 (1.44, 2.00)	Non-overweight/No Abdominal Obesity	1.00 (1.00, 1.00)
	-	-	-	Overweight/No Abdominal Obesity	1.79 (1.50, 2.15)
		-		Non-overweight/Abdominal Obesity	2.42 (1.60, 3.66)
	-	I	:	Overweight/Abdominal Obesity	2.98 (2.51, 3.53)

ALSK LACUUT	Set A Models, 1 ( controlling for abdo	Set A Models, 1 (overweight only) and 2 (overweight controlling for abdominal obesity)* Odds Ratio (95% $\rm CD^{\it C}$	l 2 (overweight i Ratio (95% CI) <sup>c</sup>	Set B Models* Odds Ratio (95% CI) Overweight <sup>c</sup> Interacted with Abdominal Obesity <sup>c</sup>	5% CI) iinal Obesity <sup>c</sup>
		(Set A 1)	(Set A 2)		(Set B)
Inflammation <i>b</i>	Overweight	2.26 (1.93, 2.64)	1.81 (1.51, 2.17)	Non-overweight/No Abdominal Obesity	1.00 (1.00, 1.00)
	Abdominal Obesity	;	1.53 (1.27, 1.85)	Overweight/No Abdominal Obesity	1.55 (1.15, 2.09)
	-	;	1	Non-overweight/Abdominal Obesity	1.36 (1.05, 1.76)
	-	;	1	Overweight/Abdominal Obesity	2.72 (2.26, 3.26)
Pre-/hypertension <sup>b</sup>	Overweight	2.45 (2.10, 2.84)	1.93 (1.62, 2.29)	Non-overweight/No Abdominal Obesity	1.00 (1.00, 1.00)
	Abdominal Obesity	;	1.74 (1.47, 2.07)	Overweight/No Abdominal Obesity	1.80 (1.37, 2.36)
	1	;	1	Non-overweight/Abdominal Obesity	1.66 (1.31, 2.09)
	-	;	1	Overweight/Abdominal Obesity	3.33 (2.81, 3.95)
High Triglycerides b	Overweight	2.63 (2.28, 3.05)v	2.09 (1.76, 2.48)	Non-overweight/No Abdominal Obesity	1.00 (1.00, 1.00)
	Abdominal Obesity	;	1.56 (1.31, 1.86)	Overweight/No Abdominal Obesity	2.00 (1.53, 2.62)
	1	;	1	Non-overweight/Abdominal Obesity	1.51 (1.18, 1.92)
	1	;	1	Overweight/Abdominal Obesity	3.25 (2.73, 3.85)
Low HDL-C <sup>b</sup>	Overweight	2.15 (1.88, 2.46)	1.73 (1.48, 2.03)	Non-overweight/No Abdominal Obesity	1.00 (1.00, 1.00)
	Abdominal Obesity	;	1.51 (1.28, 1.78)	Overweight/No Abdominal Obesity	1.53 (1.19, 1.96)
	1	;	1	Non-overweight/Abdominal Obesity	1.36 (1.09, 1.71)
	1	;	1	Overweight/Abdominal Obesity	2.58 (2.20, 3.02)
Pre-/diabetesb	Overweight	2.32 (2.03, 2.66)	1.81 (1.54, 2.13)	Non-overweight/No Abdominal Obesity	1.00 (1.00, 1.00)
	Abdominal Obesity	;	1.61 (1.37, 1.90)	Overweight/No Abdominal Obesity	1.67 (1.30, 2.16)
	-	:	:	Non-overweight/Abdominal Obesity	1.52 (1.22, 1.89)
	-	-	-	Overweight/Abdominal Obesity	2.89 (2.47, 3.39)

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nd without) control for abdominal obesity, Set beverages per week, engaging in high levels of total physical activity (across leisure, commuting, and occupational activity), urbanicity, and income

bre-diabetes or diabetes (HbA1c 5.7%) inflammation (hsCRP 3 mg/L), pre-hypertension or hypertension (SBP/DBP 130/85 mmHg), high triglycerides (150 mg/dL), low high-density lipoprotein cholesterol (<40 mg/dL, men; <50 mg/dL, women).

 $^{c}$  Overweight is defined using the recommended Asian BMI cutpoint of 23 kg/m<sup>2</sup>, with non-overweight <23 kg/m<sup>2</sup>, (17) Abdominal obesity (waist circumference: men 90 cm; women 80 cm) (13)

NOTE: Set Model A 1 and 2, interpretation: For Men, overweight alone was associated with an increased risk of inflammation of 55%, which was no longer statistically significant with the addition of abdominal ovesity into the model, whereas abdominal obesity adjuting for overweight was associated with an increased risk of 70% for inflammation

NOTE: Set Model B interpretation: For Men, the combination of overweight and abdominal obesity was associated with an increased risk of inflammation of 2.03 relative to non-overweight with no abdominal obesity