

Association between predominantly plant-based diets and iron status in Chinese adults: a cross-sectional analysis

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Abstract

Current evidence of the relationship between diets and Fe status is mostly derived from studies in developed countries with Western diets, which may not be translatable to Chinese with a predominantly plant-based diet. We extracted data that were nationally sampled from the 2009 wave of China Health and Nutrition Survey; dietary information was collected using 24-h recalls combined with a food inventory for 3 consecutive days. Blood samples were collected to quantify Fe status, and log-ferritin, transferrin receptor and Hb were used as Fe status indicators. In total, 2905 (1360 males and 1545 females) adults aged 18–50 years were included for multiple linear regression and stratified analyses. The rates of Fe deficiency and Fe-deficiency anaemia were 1.6 and 0.7% for males and 28.4 and 10.7% for females, respectively. As red meat and haem Fe consumption differed about fifteen to twenty times throughout the five groups, divided by quintiles of animal protein intake per 4.2 MJ/d, only Fe status as indicated by log-ferritin ($P=0.019$) and transferrin receptor ($P=0.024$) concentrations in males was shown to be higher as intakes of animal foods increased. Log-ferritin was positively associated with intakes of red meat ($B=0.3\%$, $P=0.01$) and haem Fe ($B=12.3\%$, $P=0.010$) in males and with intake of non-haem Fe in females ($B=2.2\%$, $P=0.024$). We conclude that diet has a very limited association with Fe status in Chinese adults consuming a traditional Chinese diet, and a predominantly plant-based diet may not be necessarily responsible for poor Fe status.

Key words: Iron-deficiency anaemia: Predominantly plant-based diets: Haem iron: Ferritin

Anaemia affects 1.62 billion of the world's population, half of which are assumed to be due to Fe deficiency (ID) according to WHO reports⁽¹⁾. ID affects all ethnic groups without exception^(2,3), and its overall prevalence is about 50% in developing countries and may reach 10% or more in developed countries⁽²⁾. ID occurs in sequentially developing stages, and the severest stage is Fe-deficiency anaemia (IDA). ID and IDA have many adverse functional consequences and socio-economic implications – for example, decreased cognitive performance, impaired physical capacity and work performance, increased morbidity from infections, and increased costs incurred by therapeutic measures⁽⁴⁾. Besides physiological (e.g. menstrual losses, repeated pregnancies) and pathological (e.g. tumours, parasitic infestations) factors⁽³⁾, insufficient Fe intake and/or low bioavailability are thought to be the main causes of ID⁽⁵⁾.

Fe is present in foods as haem or non-haem Fe. In most animal foods (such as livestock meat, poultry, fish flesh and offal), about 40% of the Fe is haem Fe^(6–8), the absorption of which is more efficient (approximately 15–35%)^(3,9) and can be enhanced by unidentified factors in meat, poultry and fish themselves^(10,11). However, in plant foods, all the Fe is

non-haem Fe, the bioavailability of which is lower (approximately 2–20%)^(3,9), whereas some constituents (e.g. ascorbic acid and possibly of carotene) contained in plant foods can enhance and other constituents (e.g. tannins, polyphenols, phytic acid and soya protein) can inhibit the absorption of non-haem Fe^(7,12–14). When consumed concurrently in the same meal, some other dietary factors can substantially enhance (e.g. livestock meat, poultry, fish and alcohol) or inhibit (e.g. eggs and Ca) non-haem Fe absorption, whereas haem Fe is less influenced^(7,12,13).

A number of cross-sectional studies in developed countries have investigated the associations between dietary factors and Fe status. Although many studies have observed that increased meat intake^(15–17) and haem Fe intake^(18–20) are associated with better Fe status (often measured by serum ferritin and Hb), a few studies, however, have found no association between meat or haem Fe intake and Fe status^(21–23). As for other dietary factors (e.g. total Fe, Ca, ascorbic acid, fruits, vegetables, dairy products, tea, coffee and alcohol) and lifestyle factors (e.g. smoking), the results of correlation analysis between Fe status and these factors have been somewhat inconsistent^(21,24,25).

Abbreviations: CHNS, China Health and Nutrition Survey; ID, iron deficiency; IDA, iron-deficiency anaemia; PAL, physical activity level.

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These enhancers and inhibitors, mostly identified from single-meal Fe absorption tests, do not always have an impact on Fe status. Studies measured over complex meals or longer periods have challenged the practical impact of single-meal studies, which may not reflect the actual Fe bioavailability of daily diet and the effects of enhancers and inhibitors^(26–28). Some studies have proved that body Fe status is also one of the determinants of dietary Fe absorption and shows an inverse relationship with both haem and non-haem Fe absorptions, and non-haem Fe absorption seems to be more responsive^(29–32). When Fe stores are low, non-haem Fe absorption is nearly as efficient as haem Fe, and when Fe stores are high it can be more completely limited^(7,29,33).

The above discoveries were mainly derived from studies based on populations or individuals in developed countries. However, dietary patterns have been identified to be largely different between developed and developing countries^(34,35). Populations of most developed countries (mainly in Europe and North America) follow a Western diet, which is characterised by higher intakes of processed meat, red meat (i.e. livestock meat), butter, high-fat dairy products, eggs and refined grains^(34,35). However, in developing countries such as China, most people adhere to a predominantly plant-based diet, containing higher quantities of grains, legumes, vegetables and fruits and lower quantities of animal foods^(34,35). In 2005, the consumption of livestock products was about 82.1 and 30.9 kg/person per year in developed and developing countries, respectively, according to FAO reports⁽³⁶⁾. In view of this difference, it should be questioned whether these findings on dietary and lifestyle factors related to Fe status from developed countries hold in populations of China and other developing countries as well.

Accordingly, using data from the China Health and Nutrition Survey (CHNS)⁽³⁷⁾, a nationally sampled population in China, this study aimed to determine the prevalence rates of ID and IDA in Chinese adults, and to investigate the associations between dietary, lifestyle and demographic factors and Fe status in males *v.* females aged 18–50 years, the least and most vulnerable populations to ID, respectively^(1,4), who consume a traditional Chinese and predominantly plant-based diet.

Methods

Study population

We extracted data from the CHNS, a multipurpose, longitudinal survey in China designed to examine how the social and economic transformation has affected a wide array of nutrition and health-related outcomes^(37,38). From 1989 to 2011, a total of nine rounds of the CHNS have covered twelve provinces that vary in geography, demography, economic development and public resources⁽³⁸⁾. A multistage, random cluster approach was used to draw the sample surveyed in each of the provinces. All participants provided their written informed consent.

In 2009, a total of 8641 adults (aged 18–99 years) had their blood samples collected and assessed as part of the CHNS. Participants were excluded from analyses if they had missing information (Fe status, dietary data, demographic data), had an infection (defined by high-sensitivity C-reactive protein (CRP) ≥ 3 mg/l⁽³⁹⁾) or abnormal Fe metabolism. Participants with

a history of diseases including diabetes, hypertension, myocardial infarction, stroke, asthma, goitre, and angular cheilosis were excluded. In addition, participants were excluded if they were blind, had physical disabilities, or were pregnant or lactating. On the basis of dietary and biomarker information, participants were further excluded if they had abnormal energy intakes (<2929 or $>20\,920$ kJ/d), excessive total Fe intakes (>60 mg/d) or abnormal Fe status readings (serum ferritin >1000 ng/l, Hb <80 or >200 g/l, transferrin <1.2 or >4.6 g/l, transferrin receptor >3.7 mg/l). Therefore, a sample of 5379 subjects was used to calculate the prevalence of anaemia, ID and IDA. Next, a sample of 4827 subjects (aged 18–94 years) was used for the analysis after excluding participants with moderate and severe anaemia (defined by Hb <100 g/l^(40,41)) and those being anaemic but with normal Fe storage (i.e. Hb <130 g/l but ferritin ≥ 15 ng/ml for males and Hb <120 g/l but ferritin ≥ 15 ng/ml for females⁽⁴⁾). For the regression and stratified analyses, participants aged >50 years were excluded, leading to a sample size of 2905 participants (aged 18–50 years).

Assessment of dietary intake

The measurement of dietary intake in CHNS is a combination of two parts: an individual-level three consecutive 24-h dietary recalls and a household-level food inventory weighing performed over the same 3 d (2 weekdays and 1 weekend day), which were randomly allocated from Monday to Sunday⁽⁴²⁾.

The three consecutive 24-h dietary recalls were administered by asking individuals to report all the foods consumed away from home each day, and the same daily interview was used to collect at-home individual consumption⁽⁴²⁾. Some important instructions were provided for interviewers to promote the completeness of the recalls: (a) before the survey, interviewers were required to be acquainted with the containers and food information in the family or region surveyed; (b) during the survey, for each day, enquiries were to be in accordance with mealtime order; (c) for each meal, enquiries were to be conducted following a certain food order (staple food, non-staple food, beverages, fruits and other types of food) to help respondents in recalling; (d) to remind respondents the foods easily overlooked, such as snacks between meals; (e) to remind about foods consumed away from home; and (f) the three-dimensional food models or two-dimensional photographs were recommended for food weight estimation⁽⁴³⁾.

Household food consumption was determined by examining changes in food inventory from the beginning to the end of each day, in combination with a weighing and measuring technique. All foods and condiments in home inventory, purchased from markets, picked from gardens and food waste were weighed and recorded. At the end of the survey, all remaining foods were again weighed and recorded. Individual average daily dietary intake was calculated from the household consumption according to the number of meals and days^(42,44).

The collection of both household and individual dietary intakes made it possible to check the quality of data collection by comparing the two and by correcting for errors. Thus, each individual's average daily dietary intake, calculated from the



household survey, was compared with his or her dietary intake on the basis of 24-h recall data. Where significant discrepancies were found, the household and the individual in question were revisited and asked about their food consumption to resolve these discrepancies. All field workers were trained nutritionists who were professionally engaged in nutrition work in their own counties and had participated in other national surveys. Almost all interviewers were graduates of post-secondary schools; many of them had 4-year degrees. In addition, 3 d of specific training in collection of dietary data was provided for this survey⁽⁴²⁾.

We combined the individual dietary recall data and the corresponding condiments data (mainly includes edible oil, sauces, vinegar, etc.) of the household food inventory to calculate the total dietary intake. The latest 2002 and 2004 Food Composition Table for China^(45,46) was utilised to calculate nutrient values for the dietary data in the present study. Haem Fe was estimated using the assumption that 40% Fe is present in animal products including livestock meat, poultry, offal and fish^(6–8). The average daily intake of the 3 consecutive days was used for each individual.

The dietary factors in our analysis included nutrient variables, food group variables, and tea, coffee and alcohol consumption. The nutrient variables including total energy, animal protein (i.e. protein of livestock meat, poultry, fish flesh, offal, dairy products and eggs), plant protein, diet fibre, retinol equivalent, vitamin B₁₂, folate, ascorbic acid, Ca, Mg, Zn, Cu, total Fe, haem Fe and non-haem Fe were selected from previous studies reporting possible relations with Fe status for our research interests. The food groups including cereals, potatoes, miscellaneous beans (including adzuki beans, mung beans, kidney beans, broad beans, etc.), vegetables, fruits, red meat (i.e. livestock meat), poultry, fish, dairy foods, eggs, soyabeans, and nuts and seeds were based on the latest 2007 *Chinese Dietary Guidelines*⁽⁴⁷⁾. White meat was defined as the sum of poultry and fish flesh. Tea, coffee and alcohol intakes were all used as dichotomous variables (1 = yes, 0 = no).

The procedures for calculating the total weight of a certain food group are as follows, with soyabean group as an example. First, convert the weights of all foods (e.g. soyabeans, black beans, tofu and soya milk) in the food group (i.e. soyabean group) to raw weights of their edible parts. Second, select a representative food (e.g. soyabeans) from the group and a representative nutrient (e.g. protein) for the chosen kind of food. Third, convert the weight of each food obtained from the first step to equivalent weight of soyabeans according to the ratio of the protein content of that food and soyabeans. Finally, the sum of the equivalent weights of all foods in this group was considered the total weight of the food group.

In order to control for energy intakes, all individual daily nutrient and food data, divided by individual daily total energy, were converted into nutrient density or food density in g/4.2 MJ per d (equivalent to g/1000 kcal per d) or mg/4.2 MJ per d or µg/4.2 MJ per d.

Assessment of iron status

Participants who signed informed consent for blood sampling provided samples. After an overnight fast (at least 8 h), blood

was drawn from an antecubital vein in the morning by trained phlebotomists using the standard protocol. Plasma and serum samples were frozen and stored at 86°C for laboratory analysis. The samples were analysed in a national central laboratory in Beijing (medical laboratory accreditation certificate ISO 15189:2007) with strict quality control^(39,48,49).

We focused on biomarkers related to Fe status: serum ferritin was assessed by RIA at North Institute of Bio-Tech, China reagents (Gamma counter XH-6020); Hb was measured by volume conductivity light-scatter method with a Beckman Coulter, USA reagents (Beckman Coulter LH750 automated analyzer); and transferrin and soluble transferrin receptor were determined by nephelometry method with a Siemens analyser, Germany reagents (Siemens BNP analyzer). High-sensitivity CRP assessed by immunoturbidimetry with a Denka Seiken analyser, Japan reagents (Hitachi 7600 automated analyzer), was used as an indicator to identify participants with inflammation (CRP ≥ 3 mg/l). According to WHO criteria⁽⁴⁾, ID was defined as serum ferritin concentrations <15 ng/ml, and anaemia was defined as Hb concentrations <130 g/l for males and <120 g/l for females. Participants with low serum ferritin combined with low Hb were considered as IDA. Mild anaemia was defined as a Hb concentration between 100 and 129 g/l in males and between 100 and 119 g/l in females^(40,41).

Other demographic and lifestyle variables

In our study, we focused on the main dependent variable – that is, Fe status – and the main independent variables – that is, dietary intakes. In addition, we included a number of other demographic and lifestyle variables: the continuous variables age, BMI and physical activity level (PAL) and the categorical variables residence area, highest education level and smoking status. BMI was calculated as weight in kilograms divided by the square of height in metres. Weight was measured using certified scales, with division value ≤0.1 kg. Measurements were carried out in the early morning and subjects were required to be fasting, hatless, barefoot, wearing only underclothes and to stand quietly in the middle of the scale, with weight of the two legs evenly distributed. The readings were recorded to the nearest 0.1 kg. Height was measured using column height gauges, with division value of 0.1 cm. Subjects were required to be hatless, barefoot, with untied hair and stand in attention, with the readings recorded to the nearest 0.1 cm. For PAL, a question addressing 'PAL involved in work' was administered, and the answers were as follows: (1) very light physical activity (working in a sitting position, e.g. office worker, watch repairer, etc.), (2) light physical activity (working in standing position, e.g. salesperson, laboratory technician, teacher, etc.), (3) moderate physical activity (student, driver, electrician, metal worker, etc.), (4) heavy physical activity (farmer, dancer, steel worker, athlete, etc.), (5) very heavy physical activity (loader, logger, miner, stonecutter, etc.). On the basis of the *Chinese Dietary Reference Intakes* (2000 edition)⁽⁵⁰⁾, PAL was quantised into multiples of BMR in our analysis: 1.3 × BMR for very light in both sexes, 1.6 and 1.5 × BMR for light, 1.7 and 1.6 × BMR for moderate, 2.1 and 1.9 × BMR for heavy, 2.4 and 2.2 × BMR for very heavy in males and females, respectively⁽⁵⁰⁾. Residence

area was divided into 1 = urban and 2 = rural. Highest education level was classified into low (1 = primary school and lower), middle (2 = lower middle school, upper middle school and technical or vocational school) and high (3 = college, university and higher). Smoking status was also categorised as yes (1 = former or current smoker) or no (0 = never smoker). Variables with too many missing values such as marital status, family and individual income were not included.

Statistical analysis

Statistical analyses were conducted using SPSS version 20 for WINDOWS. Ferritin was log transformed (log-ferritin) to address the lack of normality of this variable. In descriptive analyses, mean values and standard deviations were calculated for continuous variables and percentages were calculated for categorical variables. Independent samples *t* test was used to compare the means of Fe status biomarkers and dietary Fe intakes between males and females. The χ^2 Test was used for comparison of prevalence rates between males and females. We divided the population into five groups according to quintiles of animal protein intake per 4.2 MJ/d in males and females, respectively. One-way ANOVA with a test for linear trend was used to compare the means among groups, and when the homogeneity of variance was not satisfied, the Jonckheere–Terpstra test was used to detect the linear trend among groups. χ^2 Tests with a test for linear trend were used to analyse the percentages and linear trends among groups.

Multiple linear regression models were used to investigate the relationship between dietary factors and log-ferritin, and two separate models were created for nutrients or foods both in males and females. The above-mentioned nutrient variables and food group variables were, respectively, included in the ‘nutrient model’ and the ‘food model’, whereas the demographic and lifestyle variables were included in each model to control for their potential confounding effects. The ‘STEPWISE’ method was used to screen variables in each model. For this purpose, both haem Fe and non-haem Fe intakes were included in the nutrient model, and total Fe intake was not included to avoid possible collinearity. Total Fe intake was analysed in an independent partial correlation analysis after controlling for demographic, lifestyle and nutrient variables. *P* values were two-tailed, and *P* < 0.05 was considered to be statistically significant.

Results

Prevalence rates of iron deficiency and iron-deficiency anaemia in Chinese adults

We included a total of 5379 subjects with complete data on Fe biochemical indicators, including 2494 males (with 1416 aged 18–50 years and 1078 over the age of 50 years) and 2885 females (with 1730 aged 18–50 years and 1155 over the age of 50 years). The overall prevalence rates of anaemia, ID and IDA in the pooled population aged ≥ 18 years were 13.3, 11.0 and 4.3%, respectively. The rates of groups classified by age and sex are shown in Fig. 1. Females aged 18–50 years had the highest rates, whereas males of the same age group had the lowest rates. Rates

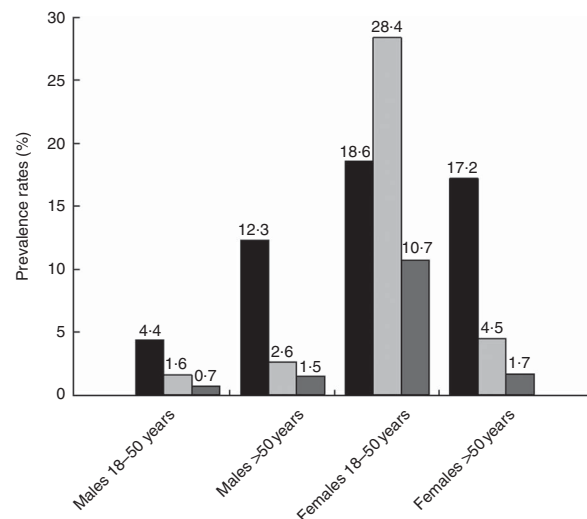


Fig. 1. Prevalence rates of anaemia (■), iron deficiency (□) and iron-deficiency anaemia (▒) in groups classified by age and sex.

of anaemia were similar between the two female groups (females 18–50 years, 18.6% *v.* females >50 years, 17.2%); however, rates of ID and IDA were vastly greater in younger females compared with older females (ID: 18–50 year, 28.4% *v.* >50 years, 4.5%; IDA: 18–50 years, 10.7% *v.* >50 years, 1.7%).

Dietary iron intake, food sources and iron status

For the analysis of dietary factors and Fe status, 1360 (46.8%) males and 1545 (53.2%) females aged 18–50 years were included in our study. Table 1 presents data on dietary intake and Fe status of males and females. The mean dietary intakes of total Fe in males and females were approximately 23.6 and 20.1 mg/d, respectively, to which haem Fe contributed approximately 4% (1.0 and 0.8 mg/d, respectively) of the total Fe intakes. Males consumed almost double the Chinese recommended nutritional intake (RNI) for Fe (12 mg/d for males), and, on average, Fe consumption of females approximately met the Chinese RNI for Fe (20 mg/d for females)⁽⁵¹⁾. The median values of serum ferritin were 123.0 and 26.7 $\mu\text{g/l}$ for males and females, respectively. Compared with females, the log-ferritin and Hb concentrations in males were much higher, whereas the concentrations of transferrin receptor were significantly lower, which indicated a higher Fe status in males compared with females.

The top ten food sources (cereals, vegetables, red meat, soyabeans, vegetable oil, white meat, algae, eggs, potatoes and miscellaneous beans) of Fe contributed to about 90% of the total dietary Fe. The ranking of their relative contributions were consistent in both males and females. The largest and second largest sources of Fe – cereals and vegetables – altogether contributed to about 60% of the total dietary Fe, followed by red meat and soyabeans, each of which contributed about 7%.

Food pattern and iron status

The population was divided into five groups according to quintiles of animal protein intakes for males and females

Table 1. Dietary iron intakes and the top ten main food sources and iron status between males and females aged 18–50 years (Mean values and percentages; mean values and standard deviations; median and 25th and 75th percentile)

	Males		Females		P*
	Mean (mg)	Contribution (%)	Mean (mg)	Contribution (%)	
Food sources of daily Fe intake					
Cereals	11.0	46.6	9.0	44.7	<0.001
Vegetables	3.4	14.2	3.0	15.0	<0.001
Red meat†	1.8	7.7	1.5	7.4	<0.001
Soyabean	1.6	7.0	1.4	6.8	0.002
Vegetable oil	0.9	4.0	0.8	4.2	<0.001
White meat‡	0.8	3.3	0.6	2.9	0.002
Algae	0.7	2.8	0.6	2.9	0.481
Eggs	0.6	2.5	0.6	2.9	0.528
Potatoes	0.3	1.2	0.3	1.4	0.451
Miscellaneous beans	0.3	1.1	0.3	1.6	0.463
	Mean	SD	Mean	SD	P
Daily Fe intakes (/d)					
Total Fe (mg)	23.6	8.0	20.1	7.3	<0.001
Haem Fe (mg)	1.0	1.2	0.8	1.0	<0.001
Non-haem Fe (mg)	22.6	7.6	19.3	7.1	<0.001
Total Fe (mg/4.2 MJ§)	9.6	2.5	10.0	2.8	<0.001
Haem Fe (mg/4.2 MJ)	0.4	0.5	0.4	0.5	0.876
Non-haem Fe (mg/4.2 MJ)	9.2	2.3	9.6	2.6	<0.001
Fe status					
Serum ferritin (µg/l)					
Median	123.0		26.7		
25th, 75th percentile	77.8, 204.3		13.5, 48.0		
Log-ferritin	4.9	0.9	3.2	1.0	<0.001
Hb (g/l)	156.0	12.6	133.6	13.5	<0.001
Transferrin receptor (mg/l)	1.3	0.4	1.5	0.5	<0.001

* Results of the comparison of mean intakes of Fe from food sources between males and females.

† Red meat, that is, livestock meat.

‡ White meat, the sum of poultry and fish flesh.

§ mg/4.2 MJ, that is, mg/1000 kcal, which is a unit of nutrient/food density used to control for energy intakes.

separately. Tables 2 and 3 show the specific dietary and Fe status data among groups.

The dietary characteristics among groups were largely consistent among males and females. Dietary patterns throughout the five groups were consistently predominantly plant-based diets, with plant foods (including cereals, potatoes, miscellaneous beans, vegetables, fruits, algae, soyabeans, nuts and seeds, plant oil and liquor) providing about 70.1–92.1 and 68.6–92.5% of the total energy in males and females, respectively. From group 1 to 5, animal protein intakes increased about ten times, from 2.0 (SD 1.4) to 20.4 (SD 5.2) g/4.2 MJ per d in males and from 1.9 (SD 1.4) to 21.4 (SD 5.6) g/4.2 MJ per d in females, with the red meat consumption increased about fourteen to fifteen times and haem Fe consumption increased about twenty times. The group consuming more animal protein in their daily diet tended to consume more red meat, white meat, eggs and dairy products and less cereals, potatoes and miscellaneous beans. Consumption of vegetables and fruits and soyabean and nuts did not show significant differences among groups. Subjects consuming greater quantities of animal foods were significantly more likely to consume tea and coffee. Females consuming more animal foods were significantly more likely to be alcohol consumers. In males, age and BMI did not differ significantly throughout the multiple groups. However, younger

females tended to consume more animal protein, and females with higher intake of animal protein had a lower BMI. PAL, the percentages of urban area and education level were significantly higher in groups with more animal protein consumption.

Throughout the five groups, both males and females tended to consume adequate quantities of Fe as per the RNI. As would be expected, red meat and haem Fe consumption tended to be significantly greater in subjects consuming more animal protein. Yet, there were no differences in Fe status (as defined by log-ferritin and transferrin receptor) in females and the prevalence rates of ID or IDA in both sexes. Despite this, the Fe status of males (as defined by log-ferritin ($P=0.019$) and transferrin receptor ($P=0.024$)) was significantly greater in groups consuming greater amounts of animal protein. However, as the intake of animal foods increased, differences in transferrin receptor were in fact very small across the quintiles among males.

Regression and correlation analyses of potential factors with log-ferritin

Table 4 shows the outcomes of linear regressions for the whole group (1360 males and 1545 females) and the non-anaemic group (1353 males and 1408 females), respectively. Both the

Table 2. Demographic factors, lifestyle factors, food and nutrient intakes and iron status in each dietary group divided by animal protein intake in males aged 18–50 years (Mean values and standard deviations; median and 25th, 75th percentile)

	Quintiles of animal protein intake per 4.2 MJ/d										<i>P</i> _{for trend}
	1 (n 271)		2 (n 274)		3 (n 271)		4 (n 272)		5 (n 272)		
	Mean	SD	Mean	SD	Mean	SD	Mean	SD	Mean	SD	
Demographic and lifestyle factors											
Age (years)	37.2	8.5	37.0	8.5	37.5	8.7	37.2	8.5	36.2	9.8	0.751
BMI (kg/m ²)	23.2	3.1	22.7	3.1	23.0	3.3	23.3	3.3	23.0	3.1	0.790
PAL (x BMR)	1.9	0.3	1.9	0.3	1.8	0.3	1.7	0.3	1.6	0.3	<0.001
Residence area											
Urban (%)	15.9		20.8		22.5		39.7		50.4		<0.001
Education level											
Low (%)	25.6		23.0		14.0		14.7		9.6		<0.001
Middle (%)	71.5		72.3		76.4		73.2		76.7		
High (%)	2.9		4.7		9.6		12.1		13.7		
Smoking status											
Former or current (%)	80.4		76.6		71.2		68.8		75.7		0.039
Daily food intakes (g/4.2 MJ per d)											
Tea consumption (yes, %)	29.9		41.2		41.4		44.1		50.7		<0.001
Coffee consumption (yes, %)	1.1		1.8		4.9		5.6		7.8		<0.001
Alcohol consumption (yes, %)	64.2		61.3		68.3		62.7		69.5		0.189
Cereals, potatoes, miscellaneous beans	200.0	37.2	179.7	35.9	166.2	30.7	153.6	33.7	138.7	34.8	<0.001
Vegetables and fruits	158.8	133.1	152.2	104.8	148.5	107.5	143.6	70.4	161.7	83.8	0.889
Red meat*	3.6	5.4	15.4	9.4	23.3	12.2	33.1	16.5	53.9	26.1	<0.001
White meat†	1.2	3.8	6.6	9.1	13.3	13.1	22.1	18.4	41.2	30.2	<0.001
Dairy products	0.2	2.5	1.7	9.3	3.4	16.2	7.2	27.0	10.0	32.7	<0.001
Eggs	7.7	8.5	10.4	11.3	13.3	13.9	14.0	15.2	14.4	15.5	<0.001
Soyabean and nuts	8.4	12.9	8.9	12.1	8.3	10.3	9.5	12.8	7.9	11.1	0.881
Daily nutrient intakes (d)											
Total energy (kJ)	10 362.9	2758.1	10 628.6	2736.8	10 861.2	2681.1	10 442.0	2661.4	9661.3	2715.4	0.002
Plant food (%TE)	92.1	9.1	84.9	8.5	79.4	8.9	75.6	9.4	70.1	9.7	<0.001
Protein (%TE)	10.8	2.1	11.0	1.8	11.5	1.7	12.8	2.0	15.2	2.6	<0.001
Fat (%TE)	24.7	9.0	29.8	9.3	32.8	8.6	34.2	8.7	35.0	9.9	<0.001
CHO (%TE)	65.9	9.3	60.3	9.0	56.9	8.4	53.7	9.1	50.4	9.7	<0.001
Animal protein (g/4.2 MJ‡)	2.0	1.4	5.7	0.9	8.9	1.0	12.7	1.3	20.4	5.2	<0.001
Plant protein (g/4.2 MJ)	23.7	5.5	20.8	4.5	18.7	4.2	18.2	4.9	16.3	4.5	<0.001
Total Fe (mg/4.2 MJ)	9.5	2.1	9.2	2.1	9.0	2.2	9.7	2.4	10.6	3.1	<0.001
Haem Fe (mg/4.2 MJ)	0.05	0.10	0.21	0.18	0.33	0.21	0.53	0.33	0.99	0.68	<0.001
Non-haem Fe (mg/4.2 MJ)	9.4	2.1	9.0	2.1	8.7	2.2	9.2	2.3	9.6	2.7	0.236
Dietary fibre (g/4.2 MJ)	5.8	3.8	5.2	2.0	4.6	1.9	5.1	3.1	4.7	2.3	<0.001
Ascorbic acid (mg/4.2 MJ)	36.2	21.8	33.0	20.3	33.5	20.8	35.6	19.8	39.9	26.0	0.017
Ca (mg/4.2 MJ)	161.2	87.0	161.2	69.2	158.2	74.0	173.9	71.3	196.8	81.8	<0.001
Retinol equivalent (µg RE/4.2 MJ)	127.8	120.4	162.0	157.9	163.3	131.1	209.2	203.1	298.0	320.1	<0.001
Zn (mg/4.2 MJ)	4.5	1.3	4.6	0.8	4.7	0.7	5.3	0.9	6.0	1.2	<0.001
Mg (mg/4.2 MJ)	141.0	34.1	133.2	26.9	124.6	26.8	130.0	29.4	132.6	29.2	<0.001
Fe status											
Serum ferritin (µg/l)											
Median	110.0		124.7		117.2		129.3		128.3		
25th, 75th percentile	70.8, 179.5		76.8, 208.0		76.3, 191.8		80.8, 220.5		80.1, 228.2		
Log-ferritin	4.8	0.8	4.9	0.8	4.9	0.8	4.9	0.9	4.9	0.9	0.019
Transferrin receptor (mg/l)	1.3	0.5	1.4	0.5	1.3	0.5	1.3	0.4	1.3	0.4	0.024
Hb (g/l)	155.2	12.1	156.5	12.9	156.0	13.0	155.8	12.0	156.1	12.7	0.631
Fe deficiency (%)	1.1		1.1		1.5		1.8		1.5		0.513
Fe-deficiency anaemia (%)	0.7		0.7		0.4		0.4		0.4		0.735

PAL, physical activity level; TE, total energy; CHO, carbohydrates; RE, retinol equivalents.

* Red meat, that is, livestock meat.

† White meat, the sum of poultry and fish flesh.

‡ g/4.2 MJ, that is, g/1000 kcal, which is a unit of nutrient/food density used to control for energy intakes.

food and nutrient models were built separately for males and females in each group.

For the whole group, the R^2 values of food and nutrient models were 8.3 and 6.6% for males and 1.3 and 1.6% for females, respectively. Of dietary variables, dietary haem Fe

($P=0.010$) was positively associated with log-ferritin in males, and its increase of 1 mg/4.2 MJ per d was accompanied by a 12.3 (95% CI 2.9, 21.7)% increase in serum ferritin concentrations, whereas dietary non-haem Fe ($P=0.024$) was positively associated with log-ferritin in females, and its increase of

Table 3. Demographic factors, lifestyle factors, food and nutrient intakes, and iron status in each dietary group divided by animal protein intake in females aged 18–50 years

	Quintiles of animal protein intake per 4.2 MJ/d										<i>P</i> _{for trend}
	1 (n 309)		2 (n 309)		3 (n 309)		4 (n 309)		5 (n 309)		
	Mean	SD	Mean	SD	Mean	SD	Mean	SD	Mean	SD	
Demographic and lifestyle factors											
Age (years)	38.6	8.3	38.8	8.0	37.7	8.5	37.1	8.5	36.1	8.5	<0.001
BMI (kg/m ²)	22.9	3.0	23.2	3.4	22.5	3.0	22.6	3.1	22.1	3.0	<0.001
PAL (×BMR)	1.7	0.2	1.7	0.2	1.6	0.2	1.5	0.2	1.5	0.2	<0.001
Residence area											
Urban (%)	14.2		19.1		26.2		40.5		51.8		<0.001
Education level											
Low (%)	44.0		37.2		30.1		20.4		16.6		<0.001
Middle (%)	55.3		58.9		65.0		67.6		71.8		
High (%)	0.7		3.9		4.9		12.0		11.6		
Smoking status											
Former or current (%)	3.9		5.5		2.9		3.2		2.6		0.149
Daily food intakes (g/4.2 MJ per d)											
Tea consumption (yes, %)	15.9		19.7		27.6		29.8		26.9		<0.001
Coffee consumption (yes, %)	0.3		2.6		4.2		8.8		8.1		<0.001
Alcohol consumption (yes, %)	3.2		5.8		10.4		12.0		15.9		<0.001
Cereals, potatoes and miscellaneous beans	199.6	39.4	177.7	36.1	164.3	34.6	148.6	31.3	134.0	35.1	<0.001
Vegetables and fruits	174.3	117.3	167.4	90.4	166.7	108.5	179.5	96.9	187.6	104.4	0.038
Red meat*	3.7	5.8	14.4	9.8	23.0	12.7	33.6	17.2	52.2	30.5	<0.001
White meat†	0.9	3.3	6.5	9.6	13.9	14	22.3	17.8	42.1	32.7	<0.001
Dairy products	0.1	2.4	0.9	5.9	4.5	15.8	9.4	29.4	20.1	48.4	<0.001
Eggs	7.3	8.9	12.2	12.8	13.7	14.5	16.8	16.4	21.4	24	<0.001
Soyabean and nuts	8.3	14.1	8.8	11.7	8.6	12.6	9.6	13.7	8.2	10	0.807
Daily nutrient intakes (d)											
Total energy (kJ)	8520.7	2554.3	8900.6	2486.1	9007.7	2468.6	8464.7	2205	7922.4	2442.2	<0.001
Plant food (%TE)	92.5	8.5	85.0	8.8	79.5	8.7	75.1	9.5	68.6	10.5	<0.001
Protein (%TE)	10.9	2.1	11.2	2.0	11.8	2.0	13.1	2.1	15.7	2.6	<0.001
Fat (%TE)	25.2	10.7	30.3	9.9	33.1	9.5	35.4	8.9	35.8	9.5	<0.001
CHO (%TE)	66.3	10.2	60.6	9.3	57.0	8.8	53.6	8.4	50.4	9.3	<0.001
Animal protein (g/4.2 MJ‡)	1.9	1.4	5.7	1	9.1	1.1	13.2	1.4	21.4	5.6	<0.001
Plant protein (g/4.2 MJ)	24.1	5.4	21	5	19.3	5.2	18.2	4.9	16.6	4.3	<0.001
Total Fe (mg/4.2 MJ)	10	2.3	9.4	2.1	9.4	2.4	10	2.9	11.2	3.6	<0.001
Haem Fe (mg/4.2 MJ)	0.05	0.09	0.19	0.16	0.34	0.22	0.53	0.4	0.99	0.7	<0.001
Non-haem Fe (mg/4.2 MJ)	9.9	2.3	9.2	2.1	9	2.4	9.5	2.8	10.2	3.3	0.089
Dietary fibre (g/4.2 MJ)	6.3	2.6	5.6	2.3	5.2	2.3	5.3	2.6	5.3	2.9	<0.001
Ascorbic acid (mg/4.2 MJ)	39.4	26	39.4	27.6	38.8	24	43.3	25.4	430	28.1	0.006
Ca (mg/4.2 MJ)	170.4	86.6	166	66.9	172.3	77.7	190	81.4	222.3	111.3	<0.001
Retinol equivalent (µgRE/4.2 MJ)	134.4	128.5	169.4	147.5	183.7	158.7	238.6	235.3	365.9	465.9	<0.001
Zn (mg/4.2 MJ)	4.5	0.7	4.7	1.3	4.8	0.8	5.2	0.9	6.2	1.2	<0.001
Mg (mg/4.2 MJ)	146.8	35	135.5	31.6	132.6	31	133.4	32.2	139.6	33.2	0.005
Fe status											
Serum ferritin (µg/l)											
Median	26.6		24.1		25.9		27.0		28.7		
25th, 75th percentile	13.9, 45.9		12.9, 47.2		13.4, 50.1		13.3, 48.3		14.4, 50.5		
Log-ferritin	3.2	1.0	3.2	1.0	3.2	1.0	3.1	1.1	3.2	1.0	0.643
Transferrin receptor (mg/l)	1.5	0.6	1.5	0.5	1.4	0.5	1.5	0.5	1.5	0.5	0.277
Hb (g/l)	133.4	12.7	135.4	15.4	134.2	14.2	132.4	13.1	132.7	11.5	0.078
Fe deficiency (%)	27.5		29.1		28.8		29.4		28.8		0.721
Fe-deficiency anaemia (%)	10.0		5.8		10.0		11.0		7.4		0.763

PAL, physical activity level; TE, Total energy.

* Red meat, that is, livestock meat.

† White meat, the sum of poultry and fish flesh.

‡ g/4.2 MJ, that is, g/1000 kcal, which is a unit of nutrient/food density used to control for energy intakes.

1 mg/4.2 MJ per d was accompanied by a 2.2 (95 % CI 0.3, 4.1) % increase in serum ferritin. In males, red meat ($P=0.010$), nuts and seeds ($P=0.001$) were positively related to log-ferritin, and potatoes ($P<0.001$) and Mg ($P=0.033$) were negatively related to log-ferritin. Being an alcohol consumer was positively associated with log-ferritin both in food models ($P=0.011$) and

nutrient models ($P=0.015$) in females. In the partial analysis, total Fe showed a significant correlation in females ($r=0.058$, $P=0.023$) but not in males ($P=0.639$). BMI was positively associated with log-ferritin in both males ($P<0.001$) and females ($P<0.05$). Age showed a positive correlation ($P<0.05$) in females but not in males.

Table 4. Multiple linear regression models for associations of foods or nutrients and other potential factors with log-ferritin (Linear regression (*B*) and 95% confidence intervals)

Variables	Whole group*			Non-anaemic group†		
	<i>B</i> ‡	95% CI	<i>P</i>	<i>B</i>	95% CI	<i>P</i>
Food model for males						
BMI (kg/m ²)	6.7	5.3, 8.1	<0.001	6.4	5.1, 7.8	<0.001
Potatoes (g/4.2 MJ per d§)	-1.7	-2.5, -0.9	<0.001	-1.5	-2.3, -0.7	<0.001
Nuts and seeds (g/4.2 MJ per d)	1.4	0.6, 2.3	0.001	1.3	0.5, 2.1	0.001
Red meat (g/4.2 MJ per d)	0.3	0.1, 0.5	0.010			
Poultry (g/4.2 MJ per d)				0.5	0.1, 0.8	0.008
Plant oil (g/4.2 MJ per d)				-0.5	-0.9, -0.1	0.026
Dairy products (g/4.2 MJ per d)				-0.2	-0.4, 0.0	0.035
<i>R</i> ² (adjust <i>R</i> ²)		0.083 (0.080)			0.088 (0.084)	
Nutrient model for males						
BMI (kg/m ²)	6.6	5.2, 8.0	<0.001	6.4	5.0, 7.7	<0.001
Haem Fe (mg/4.2 MJ per d)	12.3	2.9, 21.7	0.010	11.2	2.1, 20.3	0.016
Mg (mg/4.2 MJ per d)	-0.2	-0.3, 0.0	0.033			
<i>R</i> ² (adjust <i>R</i> ²)		0.066 (0.064)			0.063 (0.061)	
Food model for females						
BMI (kg/m ²)	1.8	0.1, 3.5	0.041			
Age (years)	0.7	0.0, 1.3	0.041	1.2	0.7, 1.7	<0.001
Alcohol (1 = yes, 0 = no)	22.1	5.0, 39.2	0.011	23.3	7.5, 39.0	0.004
<i>R</i> ² (adjust <i>R</i> ²)		0.013 (0.011)			0.019 (0.018)	
Nutrient model for females						
BMI (kg/m ²)	1.8	0.0, 3.5	0.045			
Age (years)	0.7	0.0, 1.3	0.035	1.2	0.7, 1.7	<0.001
Alcohol (1 = yes, 0 = no)	21.3	4.2, 38.4	0.015	23.3	7.5, 39.0	0.004
Non-haem Fe (mg/4.2 MJ per d)	2.2	0.3, 4.1	0.024			
<i>R</i> ² (adjust <i>R</i> ²)		0.016 (0.014)			0.019 (0.018)	

* It includes all subjects aged 18–50 years in our analysis.

† It refers to subjects who were non-anaemic of the whole group.

‡ It shows percentage increase in ferritin with 1 unit change in the variable.

§ g/4.2 MJ, that is, g/1000 kcal, which is a unit of nutrient/food density used to control for energy intakes.

|| Red meat, that is, livestock meat.

For the non-anaemic group, outcomes were generally consistent with the whole group, except that poultry ($P=0.008$), not red meat, in males was positively associated with log-ferritin, and non-haem Fe did not show any significance in females with log-ferritin. For the partial analysis in this group, total Fe still showed a marginal significance in females ($r=0.051$, $P=0.058$) but not in males ($P=0.652$).

Discussion

Dietary patterns^(3,34,35) and Fe status^(1,2) differ greatly between developed and developing countries. A number of studies have been undertaken to investigate the relationship of diets and food compositions with Fe status in developed countries with Western diets. The higher prevalence of ID in developing countries was inherently attributed to the higher intakes of plant Fe and its poor bioavailability; however, a few studies have questioned such a relation. Our study investigated Fe status and explored dietary and other potential factors related to it by using nationally sampled data in China. We found that Chinese females, who were 18–50 years old and whose total Fe intakes were about 20 mg/d, consumed diets with haem Fe at extremely different levels, from 0.5 to 8.8% of total dietary Fe, and had similar levels of ferritin, transferrin receptor and Hb as well as rates of ID and IDA. We also found a significant correlation of

Fe status with dietary haem Fe only in males with sufficient supply of dietary Fe and good Fe status. These results suggest that there is no definite association of haem Fe intakes with Fe status, and that a predominantly plant-based diet is not necessarily responsible for poor Fe status.

In China, some studies^(52,53) have provided national data on the prevalence of anaemia, whereas a few studies have reported data on the epidemiology of ID or IDA according to our literature review. Our study identified that the prevalence rates of ID approximately amounted to two and three times the corresponding rates of IDA, in accordance with the hypothesised relationship between prevalence of ID and IDA proposed in the document of WHO in 2001⁽⁴⁾. The prevalence of ID in females aged 18–50 years in China (28.4%) was higher compared with developed countries (5.2–13.2%)^(54–56), whereas the prevalence in young adult males (1.6%) was similar to that in developed countries (1–2%)^(54–56).

Compared with the median or mean intakes of Fe (8.8–13.9 mg/d)⁽²⁴⁾ among individuals in developed countries, Chinese intake far more dietary Fe (20.1–23.6 mg/d), and they have a dietary Fe density about 10 mg/4.2 MJ per d, exceeding the 7 mg/4.2 MJ per d of a typical Western diet⁽⁵⁷⁾. It seems counter-intuitive that, although the Chinese have higher intakes of total dietary Fe, yet they have higher ID rates, compared with Western countries. Possible explanations are discussed below.

A study⁽⁵⁸⁾ doubted that these dietary Fe intakes calculated on the basis of Chinese Food Composition Table and thought they might have been overestimated. However, results from the *Fourth China Total Diet Study*⁽⁵⁹⁾ estimating the dietary nutrients and contaminants through representative cooked edible foods (the mean intakes for Fe were 22.2 and 19.8 mg/d in males and females, respectively, aged 20–50 years, according to measured food Fe content) were consistent with our study and many other reports^(52,60). This gives some validation to the methods we used to quantify Fe intakes in this study.

Another explanation for high Fe intake with high ID prevalence is the quality, instead of the quantity, of dietary Fe. Considering the dietary sources of Fe, vegetables make the second largest contribution in the Chinese diet (14–15%), which is about two-fold of that in Western diets (8–9% in UK, 6.5–8.8% in New Zealand)^(21,55). Meat and meat products occupied the second position in Western diets (15–21% in UK, 19.0–22.1% in New Zealand)^(21,55), about two-fold of that in Chinese diets (10–11%). In addition, soyabean was also an important source of Fe (approximately 7% total Fe intake) in the Chinese diet, which is not commonly present in a Western-style diet. It is obvious that Fe from animal sources (haem Fe) was limited in the Chinese diet; meanwhile, a large number of plant sources provided more plant Fe, leading to a higher total Fe. Haem Fe was reported to contribute 10–15% of total Fe in Western populations consuming large amounts of animal food⁽¹²⁾, whereas in our study a much smaller proportion of haem (4%) was detected in both males and females. Owing to this fact, the higher rate of ID in Chinese females, if considered only from an ecological perspective, might be simply attributed to the low bioavailability of plant Fe⁽⁵⁾. However, the present results do not support such an explanation. In our stratified analyses, the prevalence of ID did not reduce as their haem Fe intake increased from 0.5 to 9% of total dietary Fe, a level approximating to that of Western women. In addition, from the R^2 values of each multiple linear regression model, the diet can explain a very limited portion of the ferritin variation. This hypothesis of high plant food intakes resulting in poor Fe status also could not explain the fact that Chinese males, whose dietary Fe was mostly from plant foods, have a similar ID rate as Western males.

Animal foods are the richest sources of bioavailable dietary Fe⁽⁶¹⁾. A latest systematic review⁽⁶²⁾ has investigated the association between consumption of animal flesh foods and Fe status among adults in Western countries. Findings from the review were inconsistent, but the overall conclusion was that animal flesh consumption is associated with better Fe status in developed nations, indicated by most of the positive and neutral quality studies. From this perspective, our results differed from that review, and situations were different between sexes. Red meat showed significant positive correlations in males, consistent with most association analysis studies^(16,63–65), but not in females. These results were not unprecedented and are consistent with some studies^(66,67), which observed higher ferritin concentrations or lower incidence of ID in omnivore males but not females. Moreover, our result showing a lack of relationship between animal flesh intake and Fe status in women of childbearing age was also specifically mentioned in that

review⁽⁶²⁾, which indicated that significant non-dietary factors might be major influencing factors for Fe status compared with diet for the specific sub-group.

We found that haem Fe as well as red meat did not show any benefit on Fe status in females, and several studies in different populations have shown similar findings, including populations in the UK⁽²¹⁾, young Japanese women⁽²²⁾ and healthy Danish men and women⁽⁶⁸⁾. Distinctly for females, non-haem Fe was positively related to log-ferritin in our regression analysis. This result contrasts with previous major findings, but it is not unique. A study in rural central Mexico⁽⁶⁹⁾ also reported that a better Fe status was associated with greater intakes of foods containing non-haem Fe. The similarities between findings from China and Mexico regarding Fe status and non-haem Fe consumption might reflect an adaptive change in plant Fe absorption in populations with largely plant-containing diets in developing countries. As the dietary Fe, especially non-haem Fe absorption, was inversely related to body Fe stores⁽³³⁾, non-haem Fe absorption was up-regulated in participants with low body Fe stores, suggesting that non-haem Fe can contribute more significantly to the amount of total Fe absorbed, because of enhanced bioavailability⁽²⁹⁾. This might explain why non-haem Fe was more responsive than haem Fe to differences in body Fe status in Chinese females of our study.

As for other dietary factors, alcohol consumption^(16,70) was positively and dairy products⁽⁷¹⁾ were negatively associated with log-ferritin in females and males, respectively, consistent with findings from previous studies. Mg was found to be negatively related to log-ferritin in males, which was inconsistent with previous studies^(49,72) finding that serum Mg or Mg intakes were positively associated with better Fe status. The exact mechanism of the association between Mg and Fe status is unclear at present. The most frequently mentioned enhancer ascorbic acid and inhibitors Ca, tea and coffee consumption^(7,12,14) did not show any significance in our study. From the values of R^2 of each multiple linear regression model, we can see that the diet explains a very small portion of the ferritin variation, especially in females. Those non-dietary factors not included in our analysis such as menstrual blood flow⁽²²⁾ in females might be more important in influencing factors of Fe status compared with diet.

In the 1990s, the theory that Fe absorption adapted to maintain a set point of individual homeostasis of Fe has been raised^(73,74). This adaptation was then interpreted as two main aspects: adaptation to the body's Fe status and to the dietary Fe availability over time. One aspect is reflected in the inverse relationship between Fe absorption and Fe stores^(29,30). For the other aspect, individuals tend to absorb less Fe with the high-bioavailability diet or Fe supplementation and more Fe with the low-bioavailability diet^(31,32,75). The underlying origin for the effective adaptive mechanisms for Fe was assumed to prevent both ID and Fe overload for survival^(29,30).

This adaptive control of Fe absorption may explain why diet showed little effect on Fe status in our study and why haem Fe was positively related to log-ferritin in males but not in females, which may reflect the differences in metabolic regulation responding to the dietary plant Fe between males and females. Chinese females of childbearing age, who consume large



quantities of plant foods and required more Fe, might develop great potential of adaptation to low-bioavailability Fe for their high turnover rate of body Fe, and therefore their utilisation of haem and non-haem Fe might be not that different, and the correlation between diet and Fe status may become weakened consequently. On the other hand, the situation is quite different for Chinese males. Males tend to be at greater risk of Fe overload than females and their Fe requirements are relatively low, and thus they are under less stress to adaptively improve the utilisation of non-haem Fe. Therefore, haem Fe utilisation is less regulated by the body Fe status, and dietary intakes thus are reflected in Fe storage levels in males. Our conjecture is that the adaptation in females might be higher in magnitude than males in Chinese populations, differs from other studies^(31,32) and requires further verification.

In spite of the existence of such an adaptive mechanism for regulating Fe absorption described above, ID is widespread in both developed and developing countries, and the ID rate remains high in women of childbearing age in our study. The obvious assumption may be that the enhanced Fe absorption is not sufficiently effective to prevent ID in those with high physiological Fe requirements and lower Fe intakes⁽³⁰⁾. However, our findings do not seem to support the assumption of insufficient dietary supply, as Fe storage, even at a very low level, is not associated with dietary Fe, definitely in Chinese females whose Fe intakes were maintained about 20 mg/d, implying other explanations for the higher prevalence of ID with a relatively sufficient Fe intake.

An evolutionary perspective provides a possible explanation. It is known that Fe is an essential nutrient for both humans and pathogenic microbes, and hypoferraemia, known as the secondary result of an innate immune defense, can restrict essential Fe to pathogens and inhibit pathogen proliferation^(76–78). A variety of evidences indicate that ID protects against infectious diseases and decreases the odds of death from infectious diseases⁽²⁾. Historically, with the emergence of the Agrarian civilisation in ancient China, increased crowding and poor hygiene in settlements likely increased human infestation^(79,80). It is suggested that under a selection pressure of infectious diseases, the ID phenotype survived better⁽²⁾. ID or IDA then is suggested to be an outcome of successful nutritional adaptation to infectious diseases after a long period (several millennia) of evolution, which may explain the widespread nature of ID around the world^(2,81). From this perspective, the differences in Fe status situations between developed and developing countries may be due to unbalanced advances in modern medicine (just begun a few generations ago)⁽²⁾ leading to variant adaptation pressures to infectious diseases. Although the sanitation in China has improved markedly⁽⁸²⁾ over the last few decades, there is still a big gap between China and the Western countries. For the higher rates of ID observed in our study, the relationship between environmental health conditions and Fe homeostasis need to be further explored. In general, we should take a more comprehensive view of the hazards and possible benefits (i.e. the protective effect against infection) of ID, especially when performing interventions to correct ID.

Our study has some limitations. As exposure (diet) and outcome (Fe status) are measured at the same time, cross-sectional

studies are unable to identify causal relationships. The three consecutive 24-h dietary assessment method itself has some limitations (e.g. misreporting). However, the reporting accuracy was thought to be acceptable according to a previous study⁽⁸³⁾ that evaluated the accuracy of the dietary assessment method used in the same survey. In addition, several important factors were not considered in our study, such as genetic variations⁽¹⁶⁾, information on blood donations⁽²³⁾ and menstruation in females⁽²²⁾. More well-designed studies are required to explore the relationships between these factors and Fe status in Chinese populations.

In conclusion, for Chinese females of childbearing age, dietary non-haem but not haem Fe intake is associated with Fe status in a condition of sufficient total Fe supply. For Chinese males with sufficient supply of dietary Fe and good Fe status, Fe status is related to dietary haem Fe and red meat intakes. The diet has a very limited association with Fe status in Chinese adults, indicating that a predominantly plant-based diet, as per Traditional Chinese dietary patterns, may not be entirely responsible for the poor Fe status of the Chinese population. In order to explain the high prevalence of ID in Chinese females, we need to explore other lifestyle and health factors concurrently.

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The authors declare that there are no conflicts of interest.

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