| 1 | Associations between alcohol and obesity in more than 100,000 adults in England and |
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| 2 | Scotland |
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16 Shot title: Alcohol consumption and obesity risk

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- 18 Key words: Observational study; surveys and questionnaires; overweight; alcohol drinking.

19 Abstract

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21 The objective of this cross-sectional study was to clarify the association between alcohol and 22 obesity using data from 106,182 adults in England and Scotland (46.7% male; 46.9±16.9 23 years [mean±SD]). Trained interviewers asked participants about alcohol intake. Obesity was defined as body mass index \geq 30 kg·m⁻². Potential confounders included age, sex, 24 25 smoking, physical activity, longstanding illness, psychological distress, and socioeconomic status. Compared with those who drank at least five times a week, obesity risk was 1.21 26 27 (95% confidence interval: 1.15, 1.27) in those who drank one to four times a week, 1.53 (1.43, 1.62) in those who drank one to two times a month, 1.61 (1.52, 1.71) in those who 28 drank less than once every couple of months, 1.34 (1.23, 1.47) in those who were former 29 30 drinkers, and 1.03 (0.95, 1.11) in those who were never drinkers. Compared with those who drank a harmful volume, obesity risk was 0.78 (0.68, 0.90) in those who drank within 31 32 guidelines, 0.69 (0.54, 0.88) in former drinkers, and 0.50 (0.40, 0.63) in never drinkers; And, 33 these associations were biased away from the null after adjustment for drinking volume. 34 Abstinence was associated with increased risk of obesity in women. These data suggest that 35 the association between drinking frequency and obesity is bell-shaped, with obesity risk not 36 significantly different in those who drink most often and never drinkers. Drinking volume has 37 a positive confounding effect on the association between drinking frequency and obesity, 38 which may help explain the conflicting findings of other studies.

39 Introduction

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The UK government has slashed drinking guidelines, citing concerns about cancer, heart 41 disease and other health risks ⁽¹⁾. Obesity is associated with cancer, heart disease and other 42 health risks ⁽²⁾ and the government has also raised concerns about middle-aged people 43 drinking too much and weighing too much ⁽³⁾. Observational studies of alcohol consumption 44 and obesity have produced conflicting findings ^(4; 5). There are many factors that make it 45 difficult to determine the independent influence of alcohol on obesity ⁽⁴⁾. It is particularly 46 difficult to distinguish the influence of drinking frequency from drinking volume ⁽⁴⁾. A frequent 47 48 drinking pattern may go hand in hand with lifestyle choices that decrease the risk of obesity like taking exercise and eating fruits and vegetables ^(4; 6). Conversely, a high drinking volume 49 may increase the risk of obesity ⁽⁴⁾ and the risks of morbidity and mortality ⁽⁷⁾. Socioeconomic 50 factors may also confound the association between alcohol and obesity. Lower 51 socioeconomic groups tend to make worse lifestyle choices than higher socioeconomic 52 groups in the United Kingdom^(8;9). At the same time, drinking volume tends to increase with 53 socioeconomic status ⁽⁷⁾. More research is required to clarify the link between alcohol intake 54 and obesity and to explain the conflicting findings ⁽⁴⁾. The objective of this cross-sectional 55 study was to help clarify the association between alcohol and obesity using a pooled 56 57 analysis of 11 population-based studies.

58 Methods

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60 Participants

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62 The Health Survey for England and the Scottish Health Survey are household-based surveillance studies that are described in detail elsewhere ^(10; 11). The present study included 63 64 participants from the Health Survey for England in 1994, 1997, 1998, 1999, 2003, 2004, 2006 and 2008. The present study also included participants from the Scottish Health 65 Survey in 1995, 1998 and 2003. The same organization carried out the surveys using 66 consistent methods ^(10; 11). The samples were selected using a multistage, stratified 67 68 probability design to be representative of the target populations of the corresponding 69 countries. Stratification was based on geographical areas and not on individual 70 characteristics: postcode (zip code) sectors were selected at the first stage and household 71 addresses selected at the second stage. Local research ethics committees approved all 72 aspects of each survey and all participants gave written informed consent.

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74 Independent variables

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Trained interviewers asked participants about alcohol intake ⁽¹¹⁾. The interviewers could also 76 77 ask those aged 18 to 24 years to fill in a booklet if they thought it would be difficult for them to give honest answers in front of other members of the household ⁽¹¹⁾. Participants were 78 79 asked about whether they drank alcohol nowadays. Those who said not were asked whether 80 they were always a non-drinker or whether they used to drink and had stopped. Drinkers 81 were asked, "How often have you had an alcoholic drink of any kind during the last 12 82 months?" And, "On how many days out of the last seven did you have an alcoholic drink?" Then, "Which day last week did you have the most to drink?" Drinkers were also asked to 83 describe the type or types of drink they had on that day. Total weekly alcohol units were 84 85 calculated by summing the units of each type of beverage and multiplying by the frequency.

86 In the UK, one unit is eight grammes of alcohol, which roughly corresponds to a 25 ml measure of spirits or a half-pint of beer ⁽¹²⁾; And, a 175 mL glass of wine contains two units of 87 alcohol⁽¹²⁾. Alcohol drinking frequency was derived from six frequency categories: at least 88 89 five times a week in the last 12 months; one to four times a week; one to two times a month; 90 less than once every couple of months; former drinker of alcohol; and, never drinker of 91 alcohol. Alcohol drinking volume was derived from six intake categories: harmful (>49 units 92 in men and \geq 35 units in women); hazardous (21-48 units in men and 14-34 units in women), 93 within guidelines (<21 units in men and <14 units in women); drinker, but not in last seven 94 days; former drinker of alcohol; and, never drinker of alcohol. The drinking guidelines used in 95 the present study were those that existed in the UK at the same period of time as the surveys and binge drinking was defined as drinking double or more the limit on the heaviest 96 day in past week (>8 units in men and >6 units in women) $^{(13)}$. 97

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99 Dependent variables

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Trained interviewers measured weight and height ⁽¹¹⁾, and body mass index (BMI) was 101 expressed as weight in kilograms divided by height in metres squared. Underweight was 102 defined as BMI <18.5 kg·m⁻², normal weight as BMI 18.5-24.9 kg·m⁻², overweight as BMI 25-103 <30 kg·m⁻², and obesity as BMI \ge 30 kg·m^{-2 (14)}. Weight was measured without shoes, heavy 104 105 garments, heavy jewellery, loose change, or keys. Height was measured with the participant 106 in the Frankfort plane and without shoes. Trained nurses measured waist circumference at 107 the midpoint of the iliac crest and costal margin (lower rib) and hip circumference at the widest point over the buttocks and below the iliac crest ⁽¹¹⁾. Waist and hip circumference 108 109 were measured over light clothing. Abdominal obesity was defined as waist-hip ratio >0.90 in men and >0.85 in women ⁽¹⁵⁾. 110

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114 The trained interviewers also asked about age, sex, smoking, physical activity, longstanding illness, psychological distress, occupation, and fruit and vegetable intake ⁽¹¹⁾. Participants 115 116 were asked, "Do you smoke cigarettes at all nowadays?" Those who answered yes were 117 regarded as current smokers. The guestionnaires used to assess physical activity in the Health Survey for England and Scottish Health Survey are described in detail elsewhere ⁽¹⁶⁾. 118 Participants were regarded as physically active if they reported meeting current physical 119 120 activity guidelines of at least 150 minutes per week of moderate-intensity physical activity, at least 75 minutes per week of vigorous-intensity physical activity, or equivalent combinations 121 ⁽¹⁷⁾. Longstanding illness was defined as any illness, disability or infirmity that had troubled 122 123 the respondent over a period of time or was likely to affect them over a period of time. A 124 longstanding illness was regarded as limiting if the respondent said that it limited their 125 activities in any way. The 12 item General Health Questionnaire was used to assess current mental health and a score of four or more was used to define psychological distress ^(18; 19). 126 127 Socioeconomic status was assessed using the four-group version of the Registrar General's 128 classification: professional and managerial occupations; skilled, non-manual occupations; 129 skilled manual occupations; and, routine and manual occupations. Participants in more 130 recent surveys were asked how many fruit and vegetables they ate yesterday and whether it 131 was more or less than usual.

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133 Statistical analysis

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Multinomial logistic regression was used to investigate associations between alcohol and BMI, a nominal dependent variable with more than two levels. Logistic regression was used to investigate associations between alcohol drinking and abdominal obesity, a dependent variable with two levels. The reference category for the dependent variable was normal BMI or normal waist-hip ratio. Odds ratio and 95% confidence interval were calculated for risk of

140 obesity according to alcohol drinking with the reference group being those who reported 141 drinking alcohol at least five times a week in the last 12 months. Regression models were 142 adjusted for potential confounders, including age, sex, smoking, physical activity, 143 longstanding illness, psychological distress, and socioeconomic status. Cigarette smoking 144 was categorised into five groups: never regular smoker; ex-smoker; less than ten per day; 145 ten to nineteen per day; twenty or more per day. Physical activity was also categorised into 146 two groups: meeting physical activity guidelines or not. Longstanding illness and psychological distress was also dichotomous variables: yes or no. Socioeconomic status 147 148 was categorised into the four groups of the Register General's classification. In secondary 149 analyses, we investigated associations in the subsamples of men and women and the 150 subsample that reported fruit and vegetable intake. In sensitivity analyses, we excluded 151 participants with diabetes or cardiovascular disease. All analyses were performed using 152 SPSS version 22 (IBM Inc.).

153 Results

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155 Participant characteristics

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157 The core samples of the Health Survey for England and the Scottish Health Survey are 158 designed to be representative; however, missing data may introduce bias in sub-samples. 159 Body mass index was not measured in 13,876 adults and BMI was the main cause of 160 missing data. Those excluded because of missing BMI were older than those included in the 161 present study (60 versus 47 years, p<0.001), were more often female (62 versus 53%, p<0.001), and were of lower socioeconomic status (3.9 versus 4.9% in professional and 162 163 managerial occupations, p<0.01). There were 106,182 participants in the present study 164 (46.7% male; 53.3% female; 46.9±16.9 years of age [mean±standard deviation]). Table 1 165 shows participants' characteristics according to drinking frequency. Some 18% of 166 participants reported drinking alcohol at least five times a week in the last 12 months, 45.2% 167 reported drinking alcohol one to four times a week, 13% reported drinking alcohol one to two 168 times a month, 14% reported drinking alcohol less than once every couple of months, 3.8% 169 reported being former drinkers of alcohol, and 6.4% reported never being drinkers of alcohol. 170 Those who reported drinking alcohol at least five times a week in the last 12 months 171 included a higher proportion of men, a higher proportion in professional and managerial 172 occupations, and a higher proportion of binge drinkers than those who reported drinking 173 alcohol less frequently. The proportion of obese individuals was lower in those who reported 174 drinking alcohol at least five times a week and those who reported drinking alcohol one to 175 four times a week than those who reported drinking alcohol less frequently. The proportion 176 meeting physical activity guidelines was higher in those who reported drinking alcohol at 177 least five times a week, those who reported drinking alcohol one to four times a week, and 178 those who reported drinking alcohol one to two times a month than those who reported 179 drinking alcohol less frequently. There were no obvious relationships between age and 180 drinking frequency, smoking and drinking frequency, or psychological distress and drinking

181 frequency. Abdominal obesity was present in 60.7% of men and only 26.4% of women, 182 which may explain why there was no obvious relationship between abdominal obesity and 183 drinking frequency in the whole sample. Less than 10% of participants were aged 18 to 24 184 years and were eligible to fill in a booklet if they wished; And, the removal of these 185 individuals did not change the results.

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187 Primary analyses

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189 Some 39% of participants were overweight and 21.6% obese. Figure 1, top, shows the 190 association between drinking frequency and obesity in 106,182 participants. Compared with 191 those who reported drinking alcohol at least five times a week in the last 12 months, the risk 192 of obesity rose in those who reported drinking alcohol less frequently before falling in those 193 who reported being former drinkers of alcohol and falling again in those who reported never 194 being drinkers of alcohol (a bell-shaped association). Figure 1, middle, shows the 195 association between alcohol volume and obesity in 49,073 participants. Compared with 196 those who reported drinking a harmful volume of alcohol in the last seven days, the risk of 197 obesity was not significantly different in those who reported drinking a hazardous volume of 198 alcohol in the last seven days, was lower in those who reported drinking a volume of alcohol 199 that was within the guidelines, and was lower in those who reported being former drinkers of 200 alcohol and those who reported never being drinkers of alcohol (a linear association). Figure 201 1, bottom, shows the association between drinking frequency and obesity after adjustment 202 for drinking volume. Drinking volume had a positive confounding effect on the association 203 between drinking frequency and obesity (the observed association was biased away from 204 the null) (Table S1 in the online supplement shows the numbers in each group, the odds 205 ratios, and the 95% confidence intervals). Table S2 in the online supplement shows the 206 associations of drinking frequency and drinking volume with overweight. Abdominal obesity 207 was present in 42.5% of the present sample. Figure 2 shows the association between 208 drinking frequency and abdominal obesity in 71,990 participants. Compared with those who

reported drinking alcohol at least five times a week in the last 12 months, the risk of
abdominal obesity was not significantly different in those who reported drinking one to four
times a week or one to two times a month; however, the risk of abdominal obesity was
higher in those who reported drinking less than once every couple of months, those who
reported being former drinkers of alcohol, and those who reported never being drinkers of
alcohol.

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216 Secondary analyses

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218 Table S3 in the online supplement shows the associations of drinking frequency with 219 overweight and obesity in 49,530 men and 56,526 women. In men, the association between 220 drinking frequency and obesity was bell-shaped, with obesity risk not significantly different 221 between those who drank most often and those who were ex-drinkers or had never drunk. In 222 women, compared with those who reported drinking alcohol at least five times a week in the 223 last 12 months, the risk of obesity rose in those who reported drinking one to four times a 224 week (odds ratio: 1.36; 95% CI: 1.26, 1.46), rose again in those who reported drinking 225 alcohol one to two times a month (1.89; 1.73, 2.06), and rose still in those who reported 226 drinking alcohol less than once every couple of months (2.03; 1.87, 2.20); the risk of obesity 227 fell but remained significantly different in women who reported being former drinkers of 228 alcohol (1.76; 1.56, 1.99) and women who reported never being drinkers of alcohol (1.47; 229 1.33, 1.64). Table S4 in the online supplement shows the association between drinking 230 frequency and abdominal obesity in 33,803 men and 38,187 women. The associations were 231 similar in the entire sample and the subsamples. Table S5 in the online supplement shows 232 that the associations between drinking frequency and overweight and obesity were similar with adjustment for fruit and vegetable consumption (n=48,905). Table S6 in the online 233 234 supplement shows that the associations between drinking frequency and overweigh and 235 obesity were similar after excluding those with diabetes or cardiovascular disease 236 (n=93,542).

237 Discussion

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The objective of this cross-sectional study was to help clarify the association between alcohol and obesity. The association between drinking frequency and obesity was bellshaped, with obesity risk not significantly different between those who drank most often and those who had never drunk. The association between drinking volume and obesity was linear, with obesity risk lowest in those who had never drunk. Drinking volume had a positive confounding effect on the association between drinking frequency and obesity, which may help explain the conflicting findings of other studies.

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This study has some notable strengths but also some limitations. The main strengths are the 247 248 large sample size, the detailed characterisation of drinking frequency, the detailed characterisation of drinking volume, and the adjustment for several potential confounders. 249 250 The main limitation is the cross-sectional design, where independent and dependent 251 variables are assessed at the same time. Alcohol intake, fruit and vegetable intake and 252 physical activity were self-reported. Participants were asked about their mental health in the 253 last four weeks, their drinking frequency in the last 12 months, and their drinking volume in the last seven days and we cannot exclude the possibility that those with a long-term history 254 of depression drink less alcohol ⁽²⁰⁾. Beer is carbohydrate rich and provides more energy 255 than wine per standard drink ⁽²¹⁾. No adjustment for alcohol type was made in the present 256 257 study; however, men are more likely to drink beer and women are more likely to drink wine ⁽⁴⁾; and, associations between alcohol and obesity were investigated separately in men and 258 women. Beer and spirit drinkers appear to have poorer diets than wine drinkers ⁽⁴⁾ and an 259 260 adjustment for fruit and vegetable intake was also made. No adjustment was made for 261 sleeping habit (insufficient sleep may be associated with greater alcohol consumption and excess body weight ⁽⁴⁾). Data from sub-samples of the Health Survey for England and the 262 263 Scottish Health Survey may not be representative of the target populations.

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265 Observational studies of alcohol consumption and obesity have produced conflicting 266 findings: some show a positive association, some show a negative association, and some show no association ^(4; 5). Many factors are thought to explain these conflicting findings and a 267 268 better characterisation of those more likely to be obese with alcohol has been called for ⁽⁴⁾. It 269 has been suggested that individuals who frequently drink moderate amounts of alcohol may enjoy a healthier lifestyle in general that may protect them from weight gain ⁽⁴⁾. In a sample 270 271 of 230,856 adults representative of the US population, it was found that men and women who frequently drank were far more likely to exercise vigorously than abstainers ⁽⁶⁾. Those 272 273 who drank most often in the present study were more physically active and were less likely 274 to be obese than those who drank less than once every couple of months and those who 275 were ex-drinkers. The detailed characterisation of drinking in the present study may explain 276 the conflicting findings of other studies. The association between drinking frequency and 277 obesity was bell-shaped, with obesity risk not significantly different between those who drank 278 most often and those who had never drunk. Drinking volume had a positive confounding 279 effect on the association between drinking frequency and obesity; that is, the observed 280 association was biased away from the null.

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282 Compared with women who reported drinking alcohol at least five times a week in the 283 present study, the risk of obesity was higher in women who reported being former drinkers of 284 alcohol and women who reported never being drinkers of alcohol. These data suggest that abstinence increases the risk of obesity in women. Although there is conflicting evidence ⁽²²⁾, 285 a frequent but moderate drinking pattern may be part of a healthy lifestyle ^(4; 6) and 286 287 longitudinal data from the Health Survey for England and the Scottish Health Survey suggest that there may be safe limits in relation to mortality risk ^(23; 24). For example, Perreault and 288 colleagues ⁽²⁴⁾ investigated the joint association of alcohol and physical activity with mortality 289 290 in 36,370 adults followed for 9.7±4.3 years and found that all-cause mortality risk was 291 considerably attenuated and cancer mortality risk was nearly nullified in those who met 292 physical activity guidelines. In the same study, cardiovascular disease mortality risk was

lower in occasional drinkers than never drinkers (occasional drinkers were those who
declared being drinkers but reported not drinking in the last seven days; never drinkers were
those who reported never having consumed alcohol) ⁽²⁴⁾.

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The present study may help to clarify the association between alcohol and obesity. The present study suggests that the association between drinking frequency and obesity is bellshaped, with obesity risk not significantly different between those who drink most often and those who have never drunk. The present study also suggests that drinking volume has a positive confounding effect on the association between drinking frequency and obesity. More longitudinal research is required to clarify the independent associations of drinking frequency and drinking volume with obesity.

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| 318 | |
| 319 | None. |
| 320 | |
| 321 | Authorship |
| 322 | |
| 323 | O'Donovan conceived the study. Stamatakis acquired the data and harmonised the |
| 324 | datasets. Hamer carried out the statistical analysis. All authors contributed substantially to |
| 325 | the study design, data interpretation, and writing of the manuscript. Hamer is the guarantor |
| 326 | and takes responsibility for the content of the manuscript, including the data and the |
| 327 | analysis. |

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| | Frequency of alcohol intake in last 12 months | | | | | |
|---------------|---|-----------|-----------|-----------|------------|-----------|
| | ≥5 times a | 1-4 times | 1-2 times | <1 every | Ex-drinker | Never |
| | week | a week | a month | couple of | | drinker |
| | | | | months | | |
| Age, years | 51.9±15.9 | 44.0±16.0 | 44.4±16.9 | 50.6±17.6 | 54.2±16.6 | 47.1±19.0 |
| Men, % | 60.3 | 51.0 | 37.0 | 29.6 | 42.1 | 39.2 |
| Obese, % | 18.6 | 19.4 | 24.1 | 28.6 | 27.7 | 21.6 |
| Abdominally | 51.1 | 39.8 | 36.4 | 42.1 | 54.0 | 43.0 |
| obese, % | | | | | | |
| Binge | 16.9 | 14.4 | 3.2 | 0.6 | 0 | 0 |
| drinking, % | | | | | | |
| Smoker, % | 26.8 | 27.3 | 26.6 | 27.7 | 31.6 | 16.5 |
| Physically | 22.1 | 25.6 | 21.3 | 14.3 | 13.4 | 13.7 |
| active, % | | | | | | |
| Longstanding | 45.0 | 37.6 | 41.5 | 50.6 | 63.2 | 42.3 |
| illness, % | | | | | | |
| Psychological | 13.0 | 13.1 | 14.9 | 16.9 | 22.6 | 16.0 |
| distress, % | | | | | | |
| Professional | 7.5 | 5.5 | 3.8 | 2.1 | 2.2 | 3.6 |
| occupation, | | | | | | |
| % | | | | | | |

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*Sample size is 106,182 except abdominal obesity, where sample size is 71,990. Age is mean±SD. Obese is BMI \ge 30 kg·m⁻². Abdominally obese is waist-hip ratio >0.90 in men and >0.85 in women. Binge drinking is drinking double or more the limit on heaviest day in past week (>8 units in men and >6 units in women). Physically active is meeting current physical

- activity guidelines of at least 150 minutes per week of moderate-intensity physical activity, at
- least 75 minutes per week of vigorous-intensity physical activity, or equivalent combinations.
- 337 Longstanding illness is any illness, disability or infirmity that troubled the participant over a
- period of time. Psychological distress is a General Health Questionnaire score of four or
- 339 more. Professional occupation is professional or managerial occupation according to the
- 340 Register General's classification.

341 [insert figure: "Figure 1, top.tif"]

342 [insert figure: "Figure 1, middle.tif"]

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- 344

Figure 1. Associations of drinking frequency and drinking volume with obesity. Top: 345 346 Association between drinking frequency and obesity in 106,182 adults. Middle: Association 347 between drinking volume and obesity in 49,073 adults. Bottom: Association between 348 drinking frequency and obesity after adjustment for drinking volume in 48,905 adults. The 349 reference category for the dependent variable was normal BMI. Multinomial regression 350 models were adjusted for age, sex, smoking, physical activity, longstanding illness, 351 psychological distress, and socioeconomic status. Underweight individuals were removed. Normal weight was defined as BMI 18.5-24.9 kg·m⁻², obesity as BMI \ge 30 kg·m⁻², overweight 352 as BMI 25-<30 kg·m⁻², and underweight as BMI <18.5 kg·m⁻². The numbers in each group, 353 354 the odds ratios, and the 95% confidence intervals are reported in Table S1 in the online supplement. 355

356 [insert figure: "Figure 2.tif"]

| 358 | Figure 2. Association between drinking frequency and abdominal obesity in 71,990 adults. |
|-----|---|
| 359 | The reference category for the dependent variable was normal waist-hip ratio. Logistic |
| 360 | regression models were adjusted for age, sex, smoking, physical activity, longstanding |
| 361 | illness, psychological distress, and socioeconomic status. Abdominal obesity was defined as |
| 362 | a waist-hip ratio >0.90 in men and >0.85 in women. The numbers in each group, the odds |
| 363 | ratios, and the 95% confidence intervals are reported in Table S4 in the online supplement. |

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