IARC HANDBOOKS

ABSENCE OF EXCESS BODY FATNESS

VOLUME 16

This publication represents the views and expert opinions of an IARC Working Group on the Evaluation of Cancer-Preventive Interventions, which met in Lyon, 5–12 April 2016

LYON, FRANCE - 2018

IARC HANDBOOKS OF CANCER PREVENTION

International Agency for Research on Cancer



2.2.3 Cancer of the stomach

In 2012, gastric cancer, or cancer of the stomach, was the fifth most commonly diagnosed cancer worldwide, with heterogeneous geographical distribution (Jemal et al., 2014). Gastric cancer can generally be classified into two subsites: cancer of the gastric cardia, which arises from the area of the stomach adjoining the gastro-oesophageal junction, and non-cardia gastric cancer, which develops in the distal stomach and represents about 73% of all gastric cancer cases globally (Colquhoun et al., 2015). Several risk factors for gastric cardia and non-cardia cancer have been identified. For example, infection with Helicobacter pylori has been strongly associated with non-cardia gastric cancer, whereas diets rich in smoked foods, salted foods (especially fish), or pickled foods, as well as cigarette smoking, appear to increase the risk of both types of gastric cancer (Kamangar et al., 2006; IARC, 2012).

In 2001, the IARC Handbook on weight control and physical activity (IARC, 2002) reviewed the studies of cancer of the gastric cardia together with studies of oesophageal adenocarcinoma, but did not provide a separate evaluation for stomach cancer (cardia or non-cardia). Since then, numerous individual and pooled cohort studies and meta-analyses, as well as several case-control studies of anthropometric measures and risk of stomach cancer have been published. Results from studies that examined this association for gastric cancer not otherwise specified (NOS) and separately for gastric cardia and non-cardia cancers are summarized here and in Tables 2.2.3a, 2.2.3b, and 2.2.3c. Studies that had fewer than 75 incident cases or that overlapped with a more recent study, as well as those that considered gastric cardia and oesophageal cancers together, were excluded.

(a) Cohort studies

(i) Gastric cancer NOS

Since 2000, at least 20 individual cohort studies (Table 2.2.3a) and six meta-analyses or pooled analyses (Table 2.2.3c) of prospective studies have examined associations of baseline BMI with gastric cancer incidence and/or mortality. Most of the individual prospective studies showed no associations with gastric cancer incidence or mortality (Table 2.2.3a). A few studies found inconsistent evidence of either positive or negative associations (Calle et al., 2003; Samanic et al., 2006; Jee et al., 2008; Persson et al., 2008; Camargo et al., 2014).

Although three pooled analyses and one meta-analysis also showed no association between high BMI and incidence of gastric cancer (Lindkvist et al., 2013) or incidence and/ or mortality (Renehan et al., 2008; Whitlock et al., 2009; Parr et al., 2010), others were suggestive of a positive association (Yang et al., 2009; Chen et al., 2013; Lin et al., 2014). In the most recent meta-analysis of 12 prospective studies of gastric cancer incidence and mortality combined and more than 41 791 gastric cancer cases, strong associations with overweight and obesity were reported in men only, but there was no evidence of heterogeneity of results according to sex (Chen et al., 2013). The same study did not show heterogeneity in results between Asian and non-Asian populations.

No associations of weight or BMI in early adulthood, usually defined as age 18–21 years, with gastric cancer incidence or mortality were found in three studies (Fujino et al., 2007; Merry et al., 2007; Tanaka et al., 2007), or of BMI change during adulthood in relation to incidence of gastric cancer (Merry et al., 2007; Rapp et al., 2008). No prospective studies of waist circumference and total gastric cancer were identified.

(ii) Cancer of the gastric cardia

Most individual prospective studies of the association between baseline BMI (or weight) and cardia gastric cancer incidence (or incidence and mortality) showed a positive association (see Table 2.2.3a), except for four studies (Tran et al., 2005; Samanic et al., 2006; Corley et al., 2008; Steffen et al., 2015). In the large meta-analysis by Chen et al., overweight was associated with a 21% higher risk (based on six studies) and obesity was associated with an 82% higher risk (based on seven studies) compared with normal BMI (18.5–24.9 kg/m²) (Chen et al., 2013). These findings were similar to those reported in an earlier meta-analysis of three prospective studies (Yang et al., 2009).

Associations of BMI in early adulthood and adult BMI change with incidence of cardia gastric cancer were examined in only one study of mortality (Merry et al., 2007). In that study, BMI at age 20 years was not associated with risk, whereas increasing BMI from age 20 years to baseline showed a positive association ($P_{trend} = 0.02$).

Although one study showed no association between sagittal abdominal diameter and risk of gastric cardia cancer (Corley et al., 2008), in the NIH-AARP cohort a 2.2-fold higher risk for the fourth versus the first quartile of waist circumference was reported, with a significant trend (O'Doherty et al., 2012). A similar positive trend of waist circumference and gastric cardia cancer risk (incidence and mortality) was also found in the EPIC study (Steffen et al., 2015).

(iii) Non-cardia gastric cancer

Findings from cohort studies and meta-analyses of excess body weight at baseline in relation to incidence of non-cardia gastric cancer are inconsistent. Neither BMI nor weight was associated with risk in most individual prospective studies (see <u>Table 2.2.3a</u>). Similarly, several meta-analyses did not show an association between BMI and risk either (<u>Yang et al., 2009</u>; Chen et al., 2013; Lin et al., 2014). However, in the Linxian General Population Trial, a significant inverse association was reported with a relative risk of 0.68 for BMI \geq 23 kg/m² versus BMI < 20 kg/m² (Tran et al., 2005), and a significant inverse association was also reported in a Swedish cohort study ($P_{trend} < 0.01$) (Samanic et al., 2006). Conversely, one individual study suggested a positive association of BMI and/or weight and risk of non-cardia gastric cancer (O'Doherty et al., 2012).

No associations were reported in the only study of BMI in early adulthood and adult BMI change in relation to incidence of non-cardia gastric cancer (<u>Merry et al., 2007</u>), or in the three studies that examined waist circumference and risk of non-cardia gastric cancer (<u>MacInnis et al.,</u> 2006; O'Doherty et al., 2012; <u>Steffen et al., 2015</u>).

(b) Case–control studies

See Table 2.2.3b.

There were a total of 11 independent reports from case-control studies on the association of BMI with risk of gastric cancer, in China, Europe, Japan, the Republic of Korea, the USA, and Venezuela. With the exception of one hospital-based study (Kim et al., 2015), in which BMI was measured at the time of initial endoscopic diagnosis, BMI was assessed through self-reports of height and body weight, referring to either a recent period (mostly 1 year) before disease diagnosis or a period in the more distant past (e.g. at age 18 years or 20 years), or both. In addition to standard adjustments for age and sex, studies were reported with variable adjustments for further confounding factors such as smoking, alcohol consumption, family history of gastric cancer, dietary variables, or *H. pylori* infection.

With regard to gastric cardia cancer, three out of four studies showed a positive association of BMI with risk. Three studies specifically addressing non-cardia cancer showed no association of recent BMI with risk, whereas two studies reported a positive association of risk with BMI at age 20 years. With regard to overall gastric cancer – without specification by subsite – three studies showed an increase in risk with increasing BMI, one showed a decrease in risk, and two showed no significant association.

Reference Cohort Location Follow-up period	Total number of subjects Sex Incidence/ mortality	Organ site or cancer type (ICD code)	Exposure categories	Exposed cases	Relative risk (95% CI)	Covariates	Comments
Stomach not otherwi	ise specified				-		-
<u>Calle et al. (2003)</u> Cancer Prevention Study II (CPS II) USA 1982–1998	404 576 Men Mortality 495 477 Women Mortality	Stomach ICD-9: 151.0–151.9	BMI 18.5-24.9 25-29.9 30-34.9 ≥ 35 $[P_{trend}]$ BMI 18.5-24.9 25-29.9 30-34.9 ≥ 35 $[P_{trend}]$	388 455 84 18 304 134 57 13	1.00 1.01 (0.88–1.16) 1.20 (0.94–1.52) 1.94 (1.21–3.13) [0.03] 1.00 0.89 (0.72–1.09) 1.30 (0.97–1.74) 1.08 (0.61–1.89) [0.46]	Age, education level, smoking, physical activity, alcohol consumption, marital status, race, aspirin use, consumption of fat and vegetables; for women, also adjusted for HRT use	
Samanic et al. (2004) United States Veterans cohort USA 1969–1996	4 500 700 Men Incidence	Stomach ICD-9: 151	Obesity Non-obese Obese Non-obese Obese	White men: 4989 309 Black men: 2089 99	1.00 1.07 (0.95–1.20) 1.00 0.98 (0.79–1.20)	Age, calendar year	Obesity defined as discharge diagnosis of obesity: ICD-8: 277; ICD-9: 278.0

Table 2.2.3a Cohort studies of measures of body fatness and cancer of the stomach

Reference Cohort Location Follow-up period	Total number of subjects Sex Incidence/ mortality	Organ site or cancer type (ICD code)	Exposure categories	Exposed cases	Relative risk (95% CI)	Covariates	Comments
Batty et al. (2005) Whitehall study of London-based male government employees United Kingdom 1967–2002	18 403 Men Mortality	Stomach	BMI 18.5-24.9 25.0-29.9 ≥ 30 [P _{trend}]	100 81 9	1.00 1.05 (0.76–1.44) 1.23 (0.59–2.58) [0.60]	Age, employment grade, physical activity, smoking, marital status, prevalent disease, weight loss in past year, BP medication, height, skinfold thickness, systolic BP, plasma cholesterol, glucose intolerance, diabetes	
<u>Kuriyama et al.</u> (<u>2005)</u> Population-based cohort Japan 1984–1992	12 485 Men Incidence 15 054 Women Incidence	Stomach ICD-9: 151.0–151.9	BMI 18.5-24.9 25.0-27.4 275-29.9 \geq 30 [P_{trend}] BMI 18.5-24.9 25.0-27.4 27.5-29.9 \geq 30 [P_{trend}]	243 50 14 7 26 17 4	1.13 (0.53–2.41) [0.91] 1.00 1.19 (0.76–1.86) 1.80 (1.06–3.05)	Age, smoking, alcohol consumption, diet, type of health insurance; for women, also adjusted for menopausal status, parity, age at menarche, age at first pregnancy	

Reference Cohort Location Follow-up period	Total number of subjects Sex Incidence/ mortality	Organ site or cancer type (ICD code)	Exposure categories	Exposed cases	Relative risk (95% CI)	Covariates	Comments
Lindblad et al. (2005) Case–control study nested in General Practitioner Research Database United Kingdom 1994–2001	11 023 Men and women Incidence	Stomach	BMI < 20 20-24 25-29 ≥ 30 [P_{trend}]	29 217 254 98	1.05 (0.69–1.58) 1.00 1.09 (0.90–1.32) 1.21 (0.94–1.56) [0.21]	Age, sex, calendar year, smoking, alcohol consumption, reflux	
Rapp et al. (2005) VHM&PP (population-based cohort) Austria 1985–2001	67 447 Men Incidence	Stomach ICD-9: 151	BMI 18.5-24.9 25-29.9 ≥ 30 $[P_{trend}]$ BMI 18.5-24.9 25-29.9 30-34.9 ≥ 35 $[P_{trend}]$	58 75 13 56 36 20 6	1.00 1.04 (0.73–1.47) 0.72 (0.40–1.33) [0.44] 1.00 0.78 (0.51–1.20) 1.28 (0.76–2.15) 1.34 (0.57–3.13) [0.48]	Age, smoking status, occupation Age, smoking status, occupation	
Samanic et al. (2006) Swedish Construction Worker Cohort Sweden 1958–1999	362 552 Men Incidence	Stomach ICD-7: 151	BMI 18.5-24.9 25-29.9 ≥ 30 $[P_{trend}]$	666 531 84	1.00 0.87 (0.77–0.97) 0.83 (0.66–1.05) [< 0.05]	Attained age, calendar year, smoking	

Reference Cohort Location Follow-up period	Total number of subjects Sex Incidence/ mortality	Organ site or cancer type (ICD code)	Exposure categories	Exposed cases	Relative risk (95% CI)	Covariates	Comments
<u>Fujino et al. (2007)</u>	46 465	Stomach	BMI			Age, study area	
JACC cohort	Men		< 18.5	54	1.00 (0.75-1.32)		
Japan	Mortality		18.5–24	569	1.00		
1988–1997			25–29	89	0.78 (0.62-0.97)		
			≥ 30	7	1.04 (0.49–2.20)		
			Weight (kg)				
			< 55	280	1.00		
			55-62	260	0.88 (0.74–1.04)		
			≥ 63	198	0.83 (0.69–1.01)		
			Weight (kg) at age 20 yr				
			< 55	339	1.00		
			55-60	210	1.04 (0.84-1.30)		
			≥ 61	157	1.17 (0.93-1.48)		
	46 465	Stomach	BMI			Age, study area	
	Women		< 18.5	37	1.44 (1.01-2.05)	0 /	
	Mortality		18.5–24	227	1.00		
			25–29	66	0.98 (0.74-1.30)		
			≥ 30	11	1.52 (0.82–2.80)		
			Weight (kg)				
			< 47	156	1.00		
			47–54	84	0.79 (0.60-1.03)		
			≥ 55	118	1.01 (0.78–1.29)		
			Weight (kg) at age 20 yr				
			< 47	167	1.00		
			47-52	72	0.97 (0.70-1.34)		
			≥ 53	95	1.25 (0.92-1.70)		
Máchová et al.	17 218	Stomach	BMI	222 total		Age, smoking,	Nested case-
<u>(2007)</u>	Men	ICD-10: C16	18.5-24.9		1.00	hypertension,	control study,
National Cancer	Incidence		25-29.9		1.05 (0.74–1.47)	height	reporting odds
Registry			≥ 30		0.92 (0.57–1.50)		ratios
Czech Republic	20 932		BMI	156 total		Age, smoking,	
1987-2002	Women		18.5-24.9		1.00	hypertension,	
	Incidence		25-29.9		0.81 (0.51–1.27)	height	
			≥ 30		0.97 (0.60–1.57)		

Reference Cohort Location Follow-up period	Total number of subjects Sex Incidence/ mortality	Organ site or cancer type (ICD code)	Exposure categories	Exposed cases	Relative risk (95% CI)	Covariates	Comments
Merry et al. (2007) Netherlands Cohort Study The Netherlands 1986–1999	4774 Men and women Incidence	Stomach, unspecified location ICD-O-3: C16.6–16.9 Histology: 8140– 8141, 8190–8231, 8260–8263, 8310, 8430, 8480–8490, 8560, 8570–8572	BMI at baseline < 20 20-24.9 25-29.9 \geq 30 [P_{trend}] BMI at age 20 yr < 20 20-21.4 21.5-22.9 23.0-24.9 \geq 25 [P_{trend}] BMI change, age 20 yr t < 0 0-3.9 4-7.9 \geq 8 [P_{trend}]	93 67 7 26 49 40 26 12 :o baseline 16 82 45	$\begin{bmatrix} 0.33 \end{bmatrix}$ $\begin{bmatrix} 0.60 & (0.37-0.99) \\ 1.00 \\ 0.92 & (0.59-1.44) \\ 0.70 & (0.42-1.18) \\ 0.82 & (0.42-1.60) \\ \begin{bmatrix} 0.72 \end{bmatrix} \\ 0.85 & (0.47-1.55) \\ 1.00 \\ 0.85 & (0.56-1.27) \end{bmatrix}$	Age, sex, smoking, education level, history of gastric ulcer or bleeding	
Reeves et al. (2007) Million Women Study United Kingdom 1996–2005	1 222 630 Women Incidence and mortality	Stomach ICD-10: C16	BMI < 22.5 22.5-24.9 25-27.4 27.5-29.9 ≥ 30 BMI < 22.5 22.5-24.9 25-27.4 27.5-29.9 ≥ 30	Incidence: 117 121 111 76 96 Mortality: 92 82 85 64 80	$\begin{array}{c} 1.26 \ (1.05-1.51) \\ 1.00 \ (0.84-1.20) \\ 1.04 \ (0.86-1.25) \\ 1.10 \ (0.88-1.38) \\ 1.04 \ (0.84-1.27) \\ 1.47 \ (1.19-1.81) \\ 1.00 \ (0.80-1.24) \\ 1.16 \ (0.93-1.43) \\ 1.34 \ (1.05-1.71) \\ 1.24 \ (0.99-1.55) \end{array}$	Age, geographical region, SES, reproductive history, smoking status, alcohol consumption, physical activity, menopausal status, time since menopause, HRT use	

Reference Cohort Location Follow-up period	Total number of subjects Sex Incidence/ mortality	Organ site or cancer type (ICD code)	Exposure categories	Exposed cases	Relative risk (95% CI)	Covariates	Comments
Tanaka et al. (2007) Population cohort from Takayama Japan 1992–2000	13 211 Men Mortality	Stomach ICD-9: 151 ICD-10: C16	BMI at baseline < 20.3 20.3-22.2 > 22.2 [P _{trend}] BMI at age 20 yr < 20.3 20.3-22.2 > 22.2 [P _{trend}]	33	1.00 0.68 (0.34–1.33) 0.53 (0.24–1.20) [0.12] 1.00 2.53 (1.18–5.43) 1.72 (0.79–3.73) [0.76]	Age, smoking, alcohol consumption, education level, physical activity, marital status	Too few incident cases in women (results not shown)
Jee et al. (2008) Cohort from the National Health Insurance Corporation Republic of Korea 1992–2006	770 556 Men Incidence 423 273	Stomach	BMI < 20.0 20.0-22.9 23.0-24.9 25.0-29.9 \geq 30.0 [P_{trend}] BMI	1808 5602 3839 3188 131	1.04 (0.97–1.13) 1.07 (1.01–1.13) 1.00 1.09 (1.02–1.16) 1.31 (1.05–1.64) [0.50]	Age, smoking Age, smoking	
	Women Incidence	ooniden	< 20.0 20.0-22.9 23.0-24.9 25.0-29.9 ≥ 30.0 $[P_{trend}]$	524 1314 1035 1132 111	0.86 (0.75-1.00) 0.90 (0.80-1.00) 1.00 0.94 (0.84-1.05) 0.84 (0.64-1.11) [0.25]	Age, shioking	
Rapp et al. (2008) VHM&PP (population-based cohort) Austria 1985–2002	28 711 Men Incidence	Stomach ICD-10: C16	BMI change per year < -0.1 -0.1 - < 0.1 0.1 - < 0.3 ≥ 0.3 $[P_{trend}]$	11 25 20 10	0.75 (0.36–1.54) 1.00 1.18 (0.65–2.13) 1.22 (0.58–2.59) [0.49]	Age, smoking status, blood glucose, occupational group, baseline BMI	

Table 2.2.3a (co	ontinued)						
Reference Cohort Location Follow-up period	Total number of subjects Sex Incidence/ mortality	Organ site or cancer type (ICD code)	Exposure categories	Exposed cases	Relative risk (95% CI)	Covariates	Comments
<u>Rapp et al. (2008)</u> (cont.)	36 938 Women Incidence		BMI change per year < -0.1 -0.1 - < 0.1 0.1 - < 0.3 ≥ 0.3 $[P_{trend}]$	19 12 19 9	1.73 (0.82–3.63) 1.00 1.73 (0.84–3.57) 1.11 (0.46–2.65) [0.73]	Age, smoking status, blood glucose, occupational group, baseline BMI	
Sjödahl et al. (2008) Nord-Trondelag Health Study Norway 1984–2002	73 133 Men and women Incidence	Stomach, adenocarcinoma ICD-7: 151.0, 151.8, 151.9	BMI < 18.5 18.5-24.9 25-29.9 ≥ 30 [P _{trend}]	3 104 110 32	0.7 (0.1–5.2) 1.0 1.0 (0.7–1.4) 1.1 (0.7–1.8) [0.74]	Age, sex, physical activity, occupation, salt intake, smoking, alcohol consumption	
Whitlock et al. (2009) Pooled analysis of 57 cohort studies Europe and North America Follow-up varied by cohort	894 576 Men and women Mortality	Stomach ICD-9: 151	BMI, per 5 kg/m ² For BMI 15–25 For BMI 25–50 For BMI 15–50	934 651	0.86 (0.70–1.05) 1.11 (0.94–1.32) 0.98 (0.90–1.07)	Study, sex, age, smoking	
Parr et al. (2010) Pooled analysis of 39 cohort studies Asia, Australia, and New Zealand 1961–1999, median follow-up 4 yr	326 387 Men and women Mortality	Stomach ICD-9: 151 ICD-10: C16	BMI 12-<18.5 18.5-24.9 25-29.9 ≥ 30 $[P_{trend}]$	NR	1.19 (0.87–1.62) 1.00 1.05 (0.88–1.25) 1.04 (0.67–1.63) [0.66]	Age, sex, smoking	
Chen et al. (2012) Population-based cohort of men China 1990–2006	142 214 Men Mortality	Stomach	BMI 15-23.5 23.5-35	757 198	0.74 (0.59–0.94) 0.96 (0.61–1.49)	Age, area, smoking, alcohol consumption, education level	

Reference Cohort Location Follow-up period	Total number of subjects Sex Incidence/ mortality	Organ site or cancer type (ICD code)	Exposure categories	Exposed cases	Relative risk (95% CI)	Covariates	Comments
Lindkvist et al. (2013) Metabolic Syndrome and Cancer Project (Me-Can) pooled analysis of	289 866 Men Incidence	Stomach ICD-7: 151	BMI, quintiles Q1 Q2 Q3 Q4 Q5 [P _{trend}]	157 134 154 197 186	1.00 0.79 (0.62–0.99) 0.84 (0.67–1.05) 1.02 (0.83–1.26) 1.00 (0.80–1.24) [0.26]	Smoking, age, study cohort, year of birth	Ranges of BMI quintiles not specified
prospective cohorts Austria, Norway, and Sweden 1972–2006, follow- up varied by cohort	288 834 Women Incidence	Stomach ICD-7: 151	BMI, quintiles Q1 Q2 Q3 Q4 Q5 $[P_{trend}]$	59 65 63 104 91	1.00 0.92 (0.65–1.31) 0.73 (0.51–1.05) 1.01 (0.72–1.40) 0.85 (0.61–1.20) [0.68]	Smoking, age, study cohort, year of birth	Ranges of BMI quintiles not specified
Bhaskaran et al. (2014) Population-based cohort: Clinical Practice Research Datalink United Kingdom 1987–2012	5 243 978 Incidence	Stomach ICD-10: C16	BMI per 5 kg/m ² increase [<i>P</i> _{trend}]	3337 total	1.03 (0.98–1.09) [0.16]	Age, sex, diabetes, smoking, alcohol consumption, SES, calendar year	Stronger association in non-smokers
Camargo et al. (2014) NIH-AARP cohort USA 1995–2006	483 700 Men and women Incidence	Stomach ICD-10: C16.0–16.9	BMI 18.5-24.9 25-29.9 30-34.9 ≥ 35 Weight, tertiles T1 T2 T3	1000 total	1.00 1.05 (0.90-1.22) 1.40 (1.16-1.68) 1.57 (1.21-2.04) 1.00 1.00 (0.86-1.17) 1.18 (1.01-1.38)	Age, sex, education level, cigarette smoking	

Reference Cohort Location Follow-up period	Total number of subjects Sex Incidence/ mortality	Organ site or cancer type (ICD code)	Exposure categories	Exposed cases	Relative risk (95% CI)	Covariates	Comments
Gastric cardia							
Samanic et al. (2004) United States Veterans cohort USA 1969–1996	4 500 700 Men Incidence	Gastric cardia ICD-9: 151.0	Obesity Non-obese Obese	White men: 841 72	1.00 1.38 (1.09–1.77)	Age, calendar year	Obesity defined as discharge diagnosis of obesity: ICD-8: 277; ICD-9: 278.0 Only 5 cases were available among Black men
Lindblad et al. (2005) Case–control study nested in General Practitioner Research Database United Kingdom 1994–2001	10 195 Men and women Incidence	Gastric cardia	BMI < 20 20-24 25-29 \ge 30 $[P_{trend}]$	2 36 55 20	0.50 (0.12–2.10) 1.00 1.37 (0.89–2.10) 1.46 (0.84–2.54) [0.04]	Age, sex, calendar year, smoking, alcohol consumption, reflux	
<u>Tran et al. (2005)</u> Linxian General Population Trial China 1986–2001	29 584 Men and women Incidence	Gastric cardia	BMI < 20 20-21 22 ≥ 23 [P _{trend}]	1089 total	1.00 0.98 (0.84–1.16) 0.96 (0.81–1.13) 0.95 (0.80–1.13) [0.51]	Age, sex	
Samanic et al. (2006) Swedish Construction Worker Cohort Sweden 1958–1999	362 552 Men Incidence	Gastric cardia ICD-7: 151.0	BMI 18.5-24.9 25-29.9 ≥ 30 $[P_{trend}]$	108 105 16	1.00 1.16 (0.88–1.52) 1.09 (0.64–1.85) [0.40]	Attained age, calendar year, smoking	

Absence of excess body fatness

Reference Cohort Location Follow-up period	Total number of subjects Sex Incidence/ mortality	Organ site or cancer type (ICD code)	Exposure categories	Exposed cases	Relative risk (95% CI)	Covariates Comments
Merry et al. (2007) Netherlands Cohort Study The Netherlands 1986–1999	4774 Men and women Incidence	Gastric cardia ICD-O-3: C16.0 Histology: 8140– 8141, 8190–8231, 8260–8263, 8310, 8430, 8480–8490, 8560, 8570–8572	BMI at baseline < 20 20-24.9 25-29.9 ≥ 30 $[P_{trend}]$ BMI at age 20 yr < 20 20-21.4 21.5-22.9 23.0-24.9 ≥ 25 $[P_{trend}]$ BMI change, age 20 yr to < 0 0-3.9 4-7.9 ≥ 8 $[P_{trend}]$	2 68 76 17 21 40 39 22 16 5 baseline 10 70 45 13	0.67 (0.16-2.80) 1.00 1.32 (0.94-1.85) 2.73 (1.56-4.79) [0.002] 0.66 (0.39-1.14) 1.00 1.02 (0.65-1.60) 0.75 (0.44-1.28) 1.47 (0.81-2.70) [0.17] 0.68 (0.34-1.35) 1.00 1.22 (0.82-1.82) 2.07 (1.08-3.97) [0.02]	Age, sex
Abnet et al. (2008) NIH-AARP cohort USA 1995–2003	480 475 Men and women Incidence	Gastric cardia ICD-O-3: C16.0 Histology: "adenocarcinoma"	BMI < 18.5 18.5-24.9 25-29.9 30-34.9 ≥ 35	1 76 128 71 31	0.70 (0.10–5.06) 1.00 1.06 (0.79–1.41) 1.70 (1.22–2.36) 2.46 (1.60–3.80)	Age, sex, cigarette smoking, alcohol consumption, education level, physical activity

Table 2.2.3a (co	ontinued)						
Reference Cohort Location Follow-up period	Total number of subjects Sex Incidence/ mortality	Organ site or cancer type (ICD code)	Exposure categories	Exposed cases	Relative risk (95% CI)	Covariates	Comments
Corley et al. (2008) Nested case- control of Kaiser Permanente Multiphasic Health Check-up cohort USA 1964–1973	3150 Men and women Incidence	Gastric cardia ICD-10: C16.0 Histology: 8140–8573	BMI < 18.5 18.5–24.9 25–29.9 ≥ 30 per 1 kg/m ² increase Sagittal abdominal dian < 20 20–22.4 22.5–25 ≥ 25 per 1 cm increase	16 12 12	- 1.00 0.91 (0.55-1.53) 2.04 (0.99-4.21) 1.04 (0.98-1.09) 1.00 0.69 (0.29-1.60) 1.17 (0.49-2.84) 1.28 (0.38-4.25) 1.03 (0.95-1.11)	Age, sex, year of health check-up BMI results also adjusted for ethnicity	
O'Doherty et al. (2012) NIH-AARP cohort USA 1995–2006	218 854 Men and women Incidence	Gastric cardia ICD-10: C16.0	BMI < 18.5 18.5–24.9 25–29.9 30–34.9 \geq 35 [P_{trend}] Weight, quartiles (sex-s) Q1 Q2 Q3 Q4 [P_{trend}]	50 79 45 15 pecific) 28 46 44	2.57 (0.62–10.65) 1.00 1.15 (0.80–1.65) 2.16 (1.41–3.29) 3.67 (2.00–6.71) [< 0.01] 1.00 1.66 (1.03–2.67) 1.53 (0.93–2.51) 2.52 (1.55–4.11) [< 0.01]	Age, sex, total energy intake, antacid use, aspirin use, NSAID use, marital status, diabetes, cigarette smoking, education level, ethnicity, alcohol consumption, physical activity, diet	

Reference Cohort Location Follow-up period	Total number of subjects Sex Incidence/ mortality	Organ site or cancer type (ICD code)	Exposure categories	Exposed cases	Relative risk (95% CI)	Covariates	Comments
<u>O'Doherty et al.</u> (<u>2012)</u> (cont.)			WC, quartiles (sex-spec Q1 Q2 Q3 Q4 [P _{trend}]	30 38 51	1.00 1.32 (0.82–2.14) 1.29 (0.82–2.04) 2.22 (1.43–3.47) [< 0.01]		
<u>Camargo et al.</u> (<u>2014)</u> NIH-AARP cohort USA 1995–2006	483 700 Men and women Incidence	Gastric cardia ICD-10: C16.0	BMI 18.5-24.9 25-29.9 30-34.9 ≥ 35 Weight, tertiles T1 T2 T3	478 total	1.00 1.10 (0.87–1.38) 1.64 (1.26–2.14) 2.24 (1.58–3.17) 1.00 1.20 (0.94–1.52) 1.53 (1.21–1.92)	Age, sex, education level, cigarette smoking	
Steffen et al. (2015) EPIC cohort 10 European countries 1992–2008	391 456 Men and women Incidence/mortality	Gastric cardia ICD-10: C16.0	BMI, quintiles Q1 Q2 Q3 Q4 Q5 [P _{trend}] Weight, quintiles Q1 Q2 Q3 Q4 Q5 [P _{trend}]	33 37 43 38	1.00 1.09 (0.68–1.77) 1.37 (0.87–2.17) 1.20 (0.74–1.94) 1.17 (0.71–1.92) [0.53] 1.00 1.14 (0.71–1.84) 1.29 (0.81–2.08) 1.11 (0.68–1.83) 1.26 (0.75–2.10) [0.48]	Age, centre, sex, education level, smoking, alcohol consumption, physical activity, diet, height	Sex-specific quintiles for weight, BMI, and WC. Cut- off points not provided, only the median values for each

Table 2.2.3a(c	ontinued)						
Reference Cohort Location Follow-up period	Total number of subjects Sex Incidence/ mortality	Organ site or cancer type (ICD code)	Exposure categories	Exposed cases	Relative risk (95% CI)	Covariates	Comments
<u>Steffen et al. (2015)</u> (cont.)			WC, quintiles Q1 Q2 Q3 Q4 Q5 [P _{trend}]	22 31 40 42 45	1.00 1.20 (0.69–2.09) 1.41 (0.83–2.40) 1.52 (0.89–2.58) 1.59 (0.93–2.73) [0.06]		
Gastric non-cardia Samanic et al. (2004) United States Veterans cohort USA 1969–1996	4 500 700 Men Incidence	Gastric non-cardia ICD-9: 151.x	Obesity Non-obese Obese Non-obese Obese	White men: 4148 237 Black men: 1958 94	1.00 1.00 (0.88–1.14) 1.00 0.99 (0.80–1.22)	Age, calendar year	Obesity defined as discharge diagnosis of obesity: ICD-8: 277; ICD-9: 278.0
Lindblad et al. (2005) Case-control study nested in General Practitioner Research Database United Kingdom 1994–2001	10 327 Men and women Incidence	Gastric non-cardia	BMI < 20 20-24 25-29 \geq 30 $[P_{trend}]$	16 70 83 23	1.75 (1.00–3.08) 1.00 1.11 (0.80–1.54) 0.87 (0.54–1.41) [0.18]	Age, sex, calendar year, smoking, alcohol consumption, reflux	
Tran et al. (2005) Linxian General Population Trial China 1986–2001	29 584 Men and women Incidence	Gastric non-cardia	BMI < 20 20-21 22 ≥ 23 $[P_{trend}]$	363 total	1.00 1.00 (0.76–1.32) 0.91 (0.68–1.20) 0.68 (0.49–0.93) [0.017]	Age, sex	

Absence of excess body fatness

Reference Cohort Location Follow-up period	Total number of subjects Sex Incidence/ mortality	Organ site or cancer type (ICD code)	Exposure of	categories	Exposed cases	Relative risk (95% CI)	Covariates	Comments
MacInnis et al. (2006) Melbourne Collaborative Cohort Study Australia 1990–2004	41 295 Men and women Incidence/mortality	Gastric non-cardia ICD-9: 151.1–151.9 ICD-10: C16.1–16.9	BMI < 25 25–29 ≥ 30 $[P_{trend}]$ Weight (kg Men: < 75 75–83 ≥ 84 $[P_{trend}]$ WC (cm) Men: < 94 94–101 ≥ 102 $[P_{trend}]$	 Women: < 62 62-70 ≥ 71 Women: < 80 80-87 ≥ 88 	68 total	$\begin{array}{c} 1.0\\ 0.5\ (0.3-1.0)\\ 1.0\ (0.5-1.8)\\ [0.76]\\ \hline \\ 1.0\\ 0.6\ (0.3-1.1)\\ 1.1\ (0.6-1.9)\\ [0.62]\\ \hline \\ 1.0\\ 0.8\ (0.4-1.4)\\ 1.1\ (0.6-2.0)\\ [0.57]\\ \hline \end{array}$	Sex, country of birth, education level, physical activity	
Samanic et al (2006) Swedish Construction Worker Cohort Sweden 1958–1999	362 552 Men Incidence	Gastric non-cardia ICD-7: 151.x	BMI 18.5-24.9 25-29.9 \geq 30 [P_{trend}]		558 426 68	1.00 0.81 (0.72–0.92) 0.78 (0.61–1.01) [< 0.01]	Attained age, calendar year, smoking	
<u>Merry et al. (2007)</u> Netherlands Cohort Study The Netherlands 1986–1999	4774 Men and women Incidence	Gastric non-cardia ICD-10: C16.1–16.5 Histology: 8140– 8141, 8190–8231, 8260–8263, 8310, 8430, 8480–8490, 8560, 8570–8572	BMI at base < 20 20-24.9 ≥ 30 [P_{trend}]	eline	12 115 99 9	1.80 (0.96-3.39) 1.00 0.97 (0.73-1.30) 0.68 (0.34-1.35) [0.13]	Age, sex, current smoking, number of cigarettes smoked per day, smoking duration, advation lavel	

Reference Cohort Location Follow-up period	Total number of subjects Sex Incidence/ mortality	Organ site or cancer type (ICD code)	Exposure categories	Exposed cases	Relative risk (95% CI)	Covariates	Comments
Merry et al. (2007)			BMI at age 20 yr				
cont.)			< 20	53	1.40 (0.91-2.15)		
			20-21.4	40	1.00		
			21.5-22.9	49	1.24 (0.80-1.91)		
			23.0-24.9	36	1.12 (0.69-1.80)		
			≥ 25	20	1.60 (0.91-2.83)		
			$[P_{\text{trend}}]$		[0.93]		
			BMI change, age 20 yr t	o baseline			
			< 0	17	0.77 (0.44-1.36)		
			0-3.9	106	1.00		
			4-7.9	61	0.85 (0.60-1.21)		
			≥ 8	14	0.86 (0.46-1.59)		
			$[P_{\text{trend}}]$		[0.77]		
<u>Abnet et al. (2008)</u>	480 475	Gastric non-cardia	BMI			Age, sex,	
NIH-AARP cohort	Men and women	ICD-O-3:	< 18.5	7	2.97 (1.38-6.39)	cigarette	
JSA	Incidence	C16.1-16.9	18.5-24.9	107	1.00	smoking,	
1995-2003		Histology:	25-29.9	123	0.80 (0.61-1.04)	alcohol	
		"adenocarcinoma"	30-34.9	61	1.08 (0.78-1.50)	consumption,	
			≥ 35	17	0.84 (0.50-1.42)	education level,	
						physical activity	
<u>Persson et al.</u>	44 453	Stomach, non-	BMI			Age, family	Similar results in
<u>2008)</u>	Women	cardia	< 20	53	1.00	history of	postmenopausa
apan Public Health	Incidence	ICD-10: C16.2-16.7	20-24.9	225	0.82 (0.61–1.11)	gastric cancer,	women only
Center-based			≥ 25	90	0.74 (0.53–1.04)	study area	
Prospective Study			$[P_{trend}]$		[0.10]		
apan		Stomach,	BMI				Similar results i
.990-2004		non-cardia,	< 20	12	1.00		postmenopausa
		differentiated	20-24.9	56	0.93 (0.50–1.74)		women only
		cancer type	≥ 25	29	1.12 (0.57–2.21)		
		ICD-10: C16.2-16.7	$[P_{\text{trend}}]$		[0.59]		

Reference Cohort Location Follow-up period	Total number of subjects Sex Incidence/ mortality	Organ site or cancer type (ICD code)		Exposed cases	Relative risk (95% CI)	Covariates	Comments
Persson et al. (2008) (cont.)		Stomach, non-cardia, undifferentiated cancer type ICD-10: C16.2-16.7	BMI < 20 20-24.9 ≥ 25 $[P_{trend}]$	37 153 52	1.00 0.79 (0.55–1.14) 0.60 (0.39–0.91) [0.01]		Similar results in postmenopausal women only
<u>Sjödahl et al. (2008)</u> Nord-Trondelag Health Study Norway 1984–2002	73 133 Men and women Incidence	Gastric non-cardia ICD-7: 151.0, 151.8, 151.9	BMI < 18.5 18.5-24.9 25-29.9 \geq 30 [P _{trend}]	2 84 92 29	1.1 (0.7–1.6)	Age, sex, physical activity, occupation, salt intake, smoking, alcohol consumption	
O'Doherty et al. (2012) NIH-AARP cohort USA 1995–2006	218 854 Men and women Incidence	Gastric non-cardia ICD-10: C16.1–16.7	BMI < 18.5 18.5-24.9 25-29.9 $30-34.9 \ge 35$ [P_{trend}] Weight, quartiles (sex-spec Q1 Q2 Q3	1 37 60 23 4 :ific) 20 35 32	1.34 (0.18–9.79) 1.00 1.32 (0.86–2.00) 1.46 (0.84–2.51) 0.99 (0.34–2.84) [0.38] 1.00 1.93 (1.10–3.38) 1.73 (0.96–3.10)	Age, sex, total energy intake, antacid use, aspirin use, NSAID use, marital status, diabetes, cigarette smoking, education level, ethnicity,	
			Q4 $[P_{trend}]$ WC, quartiles (sex-specific) Q1 Q2 Q3 Q4 $[P_{trend}]$	38	1.93 (1.05–3.54) [0.07]	alcohol consumption, physical activity, diet	

Reference Cohort Location Follow-up period	Total number of subjects Sex Incidence/ mortality	Organ site or cancer type (ICD code)	Exposure categories	Exposed cases	Relative risk (95% CI)	Covariates	Comments
<u>Camargo et al.</u> (2014) NIH-AARP cohort USA 1995–2006	483 700 Men and women Incidence	Gastric non-cardia ICD-10: C16.1–16.6	BMI 18.5-24.9 25-29.9 30-34.9 ≥ 35 Weight, tertiles T1 T2 T3	522 total	1.00 1.09 (0.83–1.43) 1.38 (0.99–1.92) 1.05 (0.61–1.82) 1.00 1.00 (0.76–1.32) 1.02 (0.77–1.34)	Age, sex, education level, cigarette smoking	
Steffen et al. (2015) EPIC cohort 10 European countries 1992–2008	391 456 Men and women Incidence/mortality	Gastric non-cardia ICD-10: C16.1–16.9	BMI, quintiles Q1 Q2 Q3 Q4 Q5 $[P_{trend}]$ Weight, quintiles Q1 Q2 Q3 Q4 Q5 $[P_{trend}]$ WC, quintiles Q1 Q2 Q3 Q4 Q5 $[P_{trend}]$	36 36 33 49 70 50 35 36 57 46 25 25 33 66 55	$\begin{array}{c} 1.00\\ 0.77\ (0.48-1.22)\\ 0.61\ (0.38-0.99)\\ 0.78\ (0.50-1.22)\\ 0.99\ (0.64-1.54)\\ [0.41]\\ 1.00\\ 0.68\ (0.44-1.06)\\ 0.67\ (0.43-1.06)\\ 1.02\ (0.68-1.55)\\ 0.84\ (0.53-1.32)\\ [0.94]\\ 1.00\\ 0.81\ (0.46-1.42)\\ 0.89\ (0.52-1.52)\\ 1.58\ (0.97-2.57)\\ 1.14\ (0.68-1.91)\\ \end{array}$	Sex, education level, smoking, alcohol consumption, physical activity, diet, height	Sex-specific quintiles for weight, BMI, and WC. Cut- off points not provided, only the median values for each

BMI, body mass index (in kg/m²); BP, blood pressure; CI, confidence interval; EPIC, European Prospective Investigation into Cancer and Nutrition; HRT, hormone replacement therapy; ICD, International Classification of Diseases; JACC, Japan Collaborative Cohort Study for Evaluation of Cancer Risk; NIH-AARP, National Institutes of Health–AARP Diet and Health Study; NSAID, non-steroidal anti-inflammatory drug; SES, socioeconomic status; VHM&PP, Vorarlberg Health Monitoring and Prevention Program; WC, waist circumference; yr, year or years

Reference Study location Period	Total number of cases Source of controls	Organ site	Exposure categories	Exposed cases	Relative risk (95% CI)	Adjustment for confounding	Comments
Stomach							
<u>Hansson et al.</u> (<u>1994)</u> Sweden 1989–1992	338 Population	Stomach	BMI at age 20 yr ≤ 21.20 21.21-22.60 22.62-24.20 ≥ 24.21 continuous ≤ 19.20 19.21-20.80 20.81-23.30 ≥ 23.21 continuous	Men: 37 40 45 84 Women: 12 18 40 28	$\begin{array}{c} 1.00\\ 1.06 \ (0.63-1.86)\\ 1.09 \ (0.66-1.82)\\ 2.16 \ (1.35-3.46)\\ 1.12 \ (1.05-1.20)\\ \hline 1.00\\ 1.39 \ (0.60-3.23)\\ 3.06 \ (1.43-6.58)\\ 2.14 \ (0.96-4.78)\\ 1.11 \ (1.02-1.21)\\ \end{array}$	Age, height	No differences were observed in the associations by age at interview (age groups: < 59 yr, 60-69 yr, and $\ge 70 \text{ yr})$ No associations were found between BMI and GC 20 yr before the interview
<u>Muñoz et al.</u> (2001) Venezuela 1991–1997	292 Population	Stomach	BMI < 18.5 18.5-25.0 > 25.0	200	11.0 (4.8–27.0) 1.0 0.3 (0.2–0.4)	Age, sex	Similar results for self-reported weight at current age. Increased risk in overweight cases with self-reported weight in childhood, adolescence, and early adulthood
<u>Inoue et al.</u> (<u>2002)</u> Japan 1988–1998	Women: 365 Population	Stomach Upper third Middle third Lower third	Current BMI < 21.08 21.08-23.56 > 23.56 < 21.08 21.08-23.56 > 23.56	72 total 155 total 127 total	1.00 1.69 (0.91–3.12) 1.07 (0.54–2.10) 1.00 0.75 (0.49–1.16) 0.80 (0.52–1.22)	Age, year, season of interview, family history of GC, smoking status, intake of raw vegetables and fish	Postmenopausal women only. <i>P</i> values for trend were non-significant among all subsites, both for current BMI and for BMI at age 20 yr
		Joner unit	< 21.08 21.08-23.56 > 23.56	12, total	1.00 1.02 (0.63–1.66) 1.16 (0.72–1.89)		

Table 2.2.3b Case-control studies of measures of body fatness and cancer of the stomach

Reference Study location Period	Total number of cases Source of controls	Organ site	Exposure categories	Exposed cases	Relative risk (95% CI)	Adjustment for confounding	Comments
Inoue et al.		·····	BMI at age 20 yr				
(2002) (cont.)		Upper third	< 21.08 21.08-23.56 > 23.56	72 total	1.00 1.33 (0.69–2.55) 1.33 (0.69–2.58)		
		Middle third	< 21.08 21.08-23.56 > 23.56	155 total	1.00 1.83 (1.14–2.94) 1.81 (1.12–2.93)		
		Lower third	< 21.08 21.08–23.56 > 23.56	127 total	1.00 0.88 (0.52–1.50) 1.31 (0.81–2.12)		
<u>Chung et al.</u> (2010) Republic of Korea 1990–2008	Men: 374 Women: 270 Hospital	Stomach	Current BMI > 35 vs ≤ 35 > 35 vs ≤ 35	Women:	1.94 (1.63–2.37) 1.65 (1.34–2.04)	Age	Study in young individuals (ages 18–45 yr)
<u>Praud et al.</u> (<u>2014)</u> Italy 1985–2007	Men: 612 Women: 387 Hospital	Stomach	BMI $< 25 \text{ vs} \ge 25$ $[P_{\text{trend}}]$ $< 25 \text{ vs} \ge 25$ $[P_{\text{trend}}]$	Men:	0.85 (0.79–0.90) [< 0.0001]	Age, sex, study, year of interview, education level, tobacco smoking, family history, total energy intake	
Kim et al. (2015) Republic of Korea 2003–2013	Men: 663 Women: 335 Hospital	Stomach	BMI measured at endoscopy < 23 23 - < 25 $\geq 25 - < 30$ ≥ 30 $[P_{trend}]$	193 175	1.00 1.25 (0.87–1.81) 1.33 (0.92–1.92) 1.27 (0.42–3.86) [0.43]	Age, smoking status, drinking status, family history of GC, <i>Helicobacter pylori</i> infection, atrophic gastritis, intestinal metaplasia, serum pepsinogen I/II ratio	No significant associations were observed when stratifying by cardia and non-cardia GC

Reference Study location Period	Total number of cases Source of controls	Organ site	Exposure categ	zories	Exposed cases	Relative risk (95% CI)	Adjustment for confounding	Comments
<u>Kim et al.</u> (2015) (cont.)			< 23 23- < 25		Women: 182 73	1.00 0.92 (0.6–1.43)		
(cont.)			$\geq 25 - < 30$ ≥ 30 $[P_{trend}]$		69 11	1.11 (0.70–1.77) 0.86 (0.33–2.26) [0.904]		
Song et al. (2015) Republic of Korea 2010–2014	1492 Population	Stomach	BMI at age 18 y 21.75 ≥ 25.3 21.75 ≥ 25.3	r	Men: Women:	1.00 1.13 (1.01–1.55) 1.00 1.25 (1.01–1.55)	Age, smoking status, alcohol drinking status, regular exercise, family history of GC, past medical history	
Gastric cardia Vaughan et al. (1995) USA (13 counties of Washington State) 1993–1990	165 Population	Gastric cardia, adenocarcinoma	BMI, percentile 1–10% 10–49% 50–89% 90–100%	25	13 52 74 25	0.8 (0.4–1.8) 1.0 1.3 (0.8–2.1) 1.6 (0.8–3.0)	Age, sex, education level, race, cigarette smoking, alcohol consumption	BMI percentiles (derived from in- person interviews) based on distribution of controls for each sex separately
<u>Chow et al.</u> (<u>1998</u>) USA 1993–1995	365 Population	Gastric cardia	< 23.12 23.12–25.08 25.09–27.31	fic) Women: < 21.95 21.95–24.12 24.13–27.43 ≥ 27.44	54 51 70 86	1.0 0.9 (0.6–1.5) 1.4 (0.9–2.1) 1.6 (1.1–2.6) [0.008]	Geographical location, age, sex, race, cigarette smoking, respondent status	BMI up to 1 yr before diagnosis for cases and date of interview for controls

Reference Study location Period	Total number of cases Source of controls	Organ site	Exposure cat	egories	Exposed cases		Relative risk (95% CI)	Adjustment for confounding	Comments
Lagergren et al. (1999) Sweden 1995–1997	262 Population	Gastric cardia	BMI 20 yr bei < 22 22–24.9 25–30 > 30 [P _{trend}] BMI at age 20 Men: < 20.7 20.7–22.1 22.2–23.7 > 23.7 [P _{trend}]	Fore interview 9 yr, quartiles (see Women: < 19.3 19.3–20.4 20.5–22.1 > 22.1		100 91 24 52 46	1.0 1.3 $(0.8-1.9)$ 2.2 $(1.4-3.4)$ 4.3 $(2.1-8.7)$ [< 0.001] 1.0 0.8 $(0.5-1.3)$ 1.2 $(0.8-1.9)$ 1.9 $(1.3-2.9)$ [< 0.001]	Age, sex, tobacco smoking, alcohol consumption, SES, reflux symptoms, intake of fruits and vegetables, energy intake, physical activity	
<u>Wu et al.</u> (<u>2001)</u> USA 1992–1997	277 Population (proxy control)	Gastric cardia	$\begin{array}{l} {\rm BMI \ at \ age \ 40} \\ {\rm Men:} \\ \leq 22 \\ > 22 - \leq 25 \\ > 25 - \leq 27 \\ > 27 \\ [P_{\rm trend}] \end{array}$	yr, quartiles (see Women: ≤ 21 > 21- ≤ 23 > 23- ≤ 25 > 25 yr, quartiles (see Women: ≤ 18 > 18- ≤ 20 > 20- ≤ 22 > 22	247 to		1.00 1.49 (1.0-2.1) 1.45 (0.9-2.3) 2.08 (1.4-3.2) [0.016] 1.00 1.13 (0.8-1.7) 1.36 (0.9-2.0) 1.71 (1.2-2.6) [0.006]	Smoking, age, sex, race, education level	
Gastric non-ca Chow et al. (1998) USA 1993–1995	ardia 365 Population	Gastric non- cardia	BMI up to 1 y Men: < 23.12 23.12–25.08 25.09–27.31 \geq 27.32 [P_{trend}]	rr before diagnos Women: < 21.95 21.95–24.12 24.13–27.43 ≥ 27.44		105 77 91	1.0 0.9 (0.6–1.4) 1.2 (0.8–1.8) 1.2 (0.8–1.8) [2.14]	Geographical location, age, sex, race, cigarette smoking, respondent status	BMI up to 1 yr before diagnosis for cases and date of interview for controls

Reference Study location Period	Total number of cases Source of controls	Organ site	Exposure ca	tegories	Exposed cases	Relative risk (95% CI)	Adjustment for confounding	Comments
Wu et al.	443	Gastric non-	BMI at age 4	0 yr, quartiles (se	ex-specific)		Smoking, age, sex,	Results did not change
<u>(2001)</u>	Population	cardia	Men:	Women:	352 total		race, education	when stratifying by
USA			≤ 22	≤ 21		1.00	level	Whites/non-Whites or
1992–1997			$> 22 - \le 25$	$> 21 - \le 23$		0.86 (0.6-1.2)		by sex
			$> 25 - \le 27$	$> 23 - \le 25$		1.00 (0.7–1.5)		
			> 27	> 25		1.10 (0.8–1.6)		
			$[P_{\text{trend}}]$			[0.57]		
			BMI at age 2	0 yr, quartiles (se	ex-specific)			
			Men:	Women:	352 total			
			≤ 20	≤ 18		1.00		
			$> 20 - \le 22$	$> 18 - \le 20$		1.21 (0.9-1.7)		
			$> 22 - \leq 24$	$> 20 - \le 22$		1.39 (1.0-2.0)		
			> 24	> 22		1.43 (1.0-2.1)		
			$[P_{\text{trend}}]$			[0.03]		

BMI, body mass index (in kg/m²); CI, confidence interval; GC, gastric cancer; SES, socioeconomic status; yr, year or years

Reference Period	Total number of studies Total number of cases	Organ site	Exposure categories	Relative risk (95% CI)	Adjustment for confounding	Comments
<u>Renehan et al.</u> (<u>2008)</u> 1996–2007	Men: 8 prospective studies 817 incident cases	Stomach	BMI per 5 kg/m² increase	0.97 (0.88–1.06)	Age (all studies) and other factors (not in all studies)	
	Women: 5 prospective studies 325 incident cases	Stomach	BMI per 5 kg/m² increase	1.04 (0.90–1.20)		
<u>Yang et al.</u> (2009) 1950–2009	12 prospective studies 9492 incident cases	Stomach	BMI Overweight and obese vs normal Obese vs normal Overweight vs normal	1.22 (1.06–1.41) 1.36 (1.21–1.54) 1.21 (1.08–1.36)	NR	No differences in risk by sex; normal, overweight, and obese are defined in most studies as BMI
	3 prospective studies	Cardia	BMI Overweight and obese vs normal Obese vs normal Overweight vs normal	1.55 (1.31–1.84) 2.06 (1.63–2.61) 1.40 (1.16–1.68)	NR	of 18.5–25, 25–29.9, and ≥ 30, respectively
	4 prospective studies	Non- cardia	BMI Overweight and obese vs normal Obese vs normal Overweight vs normal	1.18 (0.96–1.45) 1.26 (0.89–1.78) 1.16 (0.94–1.43)	NR	
<u>Chen et al.</u> (<u>2013)</u> 1994–2012	12 prospective studies 41 791 incident cases	Stomach	BMI 18.5- < 25 25-29.9 ≥ 30	1.00 1.01 (0.96–1.07) 1.06 (0.99–1.12)		Stronger associations in men in both BMI groups
	7 prospective studies	Cardia	BMI 18.5- < 25 25-29.9 ≥ 30	1.00 1.21 (1.03–1.42) 1.82 (1.32–2.49)		
	8 prospective studies	Non- cardia	BMI 18.5- < 25 25-29.9 ≥ 30	1.00 0.93 (0.82–1.05) 1.00 (0.87–1.15)		

Table 2.2.3c Meta-analyses of measures of body fatness and cancer of the stomach

Reference Period	Total number of studies Total number of cases	Organ site	Exposure categories	Relative risk (95% CI)	Adjustment for confounding	Comments
Lin et al.	13 prospective	Stomach	BMI		Age and others (not	Stronger association
<u>(2014)</u>	studies and 3		18.5- < 25	1.00	specified)	of obesity with risk in
NR	case-controls		25–29.9	1.13 (1.03–1.24)		men (5 studies) and in
	NR		≥ 30	1.04 (0.96-1.12)		non-Asian population (11
		Cardia	BMI		Age and others (not	studies)
			18.5- < 25	1.00	specified)	
			25-29.9	1.61 (1.15-2.24)	L	
			≥ 30	1.22 (1.05-1.42)		
		Non-	BMI		Age and others (not	
		cardia	18.5- < 25	1.00	specified)	
			25-29.9	0.83 (0.68-1.01)	L ·	
			≥ 30	0.94 (0.81-1.10)		

BMI, body mass index (in kg/m²); CI, confidence interval; CRC, colorectal cancer; HRT, hormone replacement therapy; IBD, inflammatory bowel disease; NR, not reported

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