#### REVIEW





# Association Between Energy and Macronutrient Intakes and Weight Change After Bariatric Surgery: a Systematic Review and Meta-analysis

Nazanin Moslehi<sup>1</sup> · Zahra Kamali<sup>2</sup> · Mahdieh Golzarand<sup>1</sup> · Fatemeh Rahimi Sakak<sup>1</sup> · Parvin Mirmiran<sup>2</sup>

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## Abstract

**Objectives** This systematic review and meta-analysis aimed to summarize the evidence on the associations of energy and macronutrient intakes (carbohydrates, fats, and proteins) with weight loss in adults after sleeve gastrectomy or gastric bypass and to determine whether these dietary characteristics of patients with suboptimal weight loss (SWL) or weight regain differ from those without these experiences.

**Methods** PubMed, Scopus, and Web of Science were searched until December 2021. Twenty-three observational studies were included.

**Results** Studies on the association of postoperative energy and macronutrients and weight loss used diverse approaches. Pooled results showed that patients with SWL consumed more energy than those with acceptable weight loss. Weight regainers consumed more energy and carbohydrates and less protein than non-regainers.

**Conclusions** Higher energy consumption is related to SWL and weight regain after surgery. Associations between macronutrients and weight outcome following bariatric surgery warrant further investigation.

Keywords Weight loss  $\cdot$  Weight regain  $\cdot$  Sleeve gastrectomy  $\cdot$  Roux-en-Y gastric bypass  $\cdot$  Carbohydrates  $\cdot$  Protein  $\cdot$  Macronutrients

#### **Key Points**

• Post-bariatric surgery dietary macronutrient and weight loss studies are limited.

• Patients with suboptimal vs. acceptable weight loss consume 203 kcal/day more energy.

• Weight regainers consumed 192 kcal/day more energy than non-regainers.

• Weight regainers consumed 25 g more carbohydrates per day than non-regainers.

☑ Nazanin Moslehi moslehinazanin@yahoo.com; moslehinazanin@sbmu.ac.ir

Parvin Mirmiran mirmiran@endocrine.ac.ir; parvin.mirmiran@gmail.com

- <sup>1</sup> Nutrition and Endocrine Research Center, Research Institute for Endocrine Sciences, Shahid Beheshti University of Medical Sciences, No. 24, Shahid Arabi St, Yemen Blvd, Chamran Exp, Tehran 19857174, Iran
- <sup>2</sup> Department of Clinical Nutrition and Dietetics, Faculty of Nutrition and Food Technology, National Nutrition and Food Technology Research Institute, Shahid Beheshti University of Medical Sciences, No. 7, Shahid Hafezi St., Farahzadi Blvd., Shahrak-E-Qods, Tehran 1981619573, Iran

## Introduction

Bariatric surgery (BS) is currently the most effective option for achieving long-term weight loss and improving obesityrelated diseases in individuals with morbid obesity [1, 2]. A meta-analysis showed a sustained weight reduction after BS for 10 years post-surgery, ranging from 45.8 to 80.9% excess weight loss (EWL) for various surgical methods [3]. However, nearly 20% of patients experienced suboptimal weight loss (SWL), defined as a percentage EWL of less than 50% in the first 12 months following BS [4]. Moreover, weight regain was seen in 5.7% of patients 2 years and 75.6% of patients 6 years after sleeve gastrectomy (SG) and in 50% of patients 2 years after Roux-en-Y gastric bypass (RYGB) [5, 6]. Therefore, the success of the surgery is not consistent among individuals, with some of them failing to achieve sufficient postoperative weight reduction (primary non-responders) or regaining most or all of the lost weight over time (secondary non-responders) [7]. These poor outcomes of BS lead to a recurrence of obesity-related comorbidities, worsen patients' quality of life, and increase the BS revision rate and healthcare cost [8, 9].

Factors that predict responses to BS are typically separated into two categories: pre- and postoperative variables. Postoperative variables are considered more significant predictors of BS success than preoperative variables [7]. Dietary factors, including dietary intake and eating behavior after surgery, are proposed as one of the main postoperative risk factors for ineffective weight loss following BS [10, 11]. Identifying the dietary factors related to weight loss success following BS is essential for establishing dietary considerations to reduce the risk of SWL and weight regain. Therefore, in this systematic review and meta-analysis, we aimed to answer the following questions: (1) what is the current evidence on the associations between energy and macronutrients including carbohydrate, fat, and protein intakes and weight loss or regain the following BS in adults? (2) Are there any differences in energy and macronutrient intakes in those with SWL after BS compared to those with acceptable weight loss (AWL)? (3) Are there any differences in energy and macronutrient intakes in weight regainers (WRs) after BS compared to non-weight regainers (non-WRs)?

## Method

The Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) statement was followed when conducting this systematic review and meta-analysis [12]. The protocol of this study was registered in the International Prospective Register for Systematic Reviews (PROSPERO) (registration number: CRD42022325140).

## Information Sources and Search Strategy

We searched PubMed/Medline, Scopus, and Web of Science from inception to December 15, 2021. The following main keywords were used: gastric sleeve, sleeve gastrectomy, gastric bypass, weight, body mass index, BMI, body composition\*, muscle mass, lean mass, carbohydrate\*, protein\* fat, fats, macronutrient\*, food\*, and eating. A full electronic search strategy for each database was provided in Supplementary Table 1. Reference lists of relevant publications were also manually searched for potentially relevant articles that were missed by the electronic search.

### **Study Selection**

After removing duplicate publications, two researchers (ZK and FRS) independently reviewed all records based on their titles and abstracts and retrieved the full texts of studies that met the inclusion criteria. The third researcher (NM) resolved discrepancies. The population, exposure, comparison, outcomes, and study design (PICOS) of this study are shown in Supplementary Table 2. Original studies that met the following inclusion criteria were included in the

study: (1) participants were over 18 years old with a history of SG or gastric bypass (GB); (2) investigated the associations of postoperative intakes of energy and macronutrients (as grams/day (g/day) or percentage of energy intake with weight outcome (either weight loss or weight regain); and (3) had observational study design. Weight loss outcome in this study included various weight loss parameters as either the absolute changes in weight (kg) and body mass index (BMI) (kg/m<sup>2</sup>) from pre-surgery or their relative changes, including %EWL, percentage of total weight loss (%TWL), and percentage of excess BMI loss (%EBMIL).

The study exclusion criteria were: (1) abstracts, reviews, conference papers, editorials, and books; (2) studies on pediatrics, adolescents, or pregnant women; (3) non-human studies; (4) studies on micronutrients or food groups intakes after BS; (5) studies on eating behaviors or food preference after BS; (6) studies assessing only preoperative dietary intakes; (7) non-English articles; (8) studies that did not report the related information; and (9) study with dietary intervention.

## **Data Extraction**

After obtaining the full text of the studies, an investigator (ZK) extracted data from eligible studies into Microsoft Excel spreadsheets, and a second investigator (NM) validated the data extraction. The following information was extracted: first author's name, country, year of publication, sample size, type of surgery, postoperative time, dietary assessment tool, dietary exposures, % of women, the definition of weight loss parameters, SWL cut-off, the definition of weight regain, weight regain cut-off, number of those with SWL/AWL, number of WRs/non-WRs, and main statistical parameters reported for the association of dietary intakes with weight outcomes.

#### **Quality Assessment**

The Newcastle–Ottawa quality assessment scale (NOS) was used to assess the quality of the studies by an investigator (MG) [13]. The NOS score goes from 0 (highest risk of bias, lowest quality) to 9 (lowest risk of bias, highest quality), with a score of  $\geq$  7 being considered high quality.

#### Data Synthesis and Analysis

The results of the included studies were quantitatively summarized. The weighted mean difference (MD) and 95% confidence interval (CI) were calculated using the random-effects model to estimate differences in energy and macronutrient consumption in patients with SWL compared to those with AWL after BS. Differences in energy and macronutrients according to weight regain were also investigated, by estimation of MD (95% CI) in dietary intakes among WRs versus non-WRs using the random-effects model. Heterogeneity between studies was assessed using the  $l^2$  statistic and Cochran's Q test; substantial heterogeneity was considered as  $l^2 > 50\%$  [14]. Egger's regression test was used to investigate publication bias. p < 0.10 was used as the significant criteria for heterogeneity and publication bias. The statistical analysis was done with Stata, version 14.2 (Stata Corp, College Station, TX).

## Results

## **Study Selection**

The process of study selection is shown in Fig. 1. The bibliographic searches in the three databases yielded 3683 unique articles. After excluding 3584 articles through screening of titles and abstracts, the full text of 99 articles was assessed for eligibility. Two articles were also found through manual reference checking of the retrieved articles [15, 16]. Ultimately, 23 studies were included in this systematic review and meta-analysis study [15–37].

#### **Study Characteristics**

Thirteen studies were on weight loss [15–24, 29, 32, 36], eight on weight regain [25–28, 31, 33, 35, 37], and two on both outcomes [30, 34]. The characteristics of studies on weight loss after BS are reported in Table 1. These studies were published between 1983 and 2022. The mean age of participants ranged between 32.9 and 48.2 years, and they underwent GB in eleven [15, 17–24, 30, 36], SG in two [29, 34], and both in two studies [16, 32]. The sample size of the studies was between 25 and 355. Five studies had follow-up time from surgery <12 months [15, 17, 19, 21, 29].

The characteristics of studies on weight regain after BS are reported in Table 2. The studies were published between 2012 and 2021. The mean age of the participants was 33.5-53.2 years; the type of surgery was GB in 6 [25, 27, 28, 30, 33, 35], SG in three [26, 31, 34], and both in one study [37]. The sample size of the studies ranged between 27 and 100, and the time since surgery for all studies was > 12 months.

#### **Quality of Studies**

The NOS score of the studies ranged from 3 to 9; four had a score  $\geq$ 7 [21, 27, 35, 36]. Details of the NOS scoring are provided in Supplementary Tables 3–5.

#### **Energy, Macronutrients, and Weight Loss**

Ten studies examined associations between energy and macronutrient intakes and weight loss as continuous variables after BS [15–19, 21, 23, 32, 34, 36]. In the first study, postoperative changes in energy and all macronutrient intakes were significantly correlated with weight change during the first 6 months after the surgery in 25 GB patients. After 1 year, however, both protein and carbohydrate intake had increased, while fat intake had plateaued. Therefore, the authors concluded that a decrease in energy intake, predominantly from fat, may be related to weight reduction after 12 months of the surgery [17]. In the other study on energy and macronutrients over 1 year and weight reduction, energy intake was significantly related to weight loss in 50 RYGB women, whereas macronutrient composition revealed no association [15]. In a prospective study on 167 individuals undergoing RYGB, daily intake of protein per kg of body weight was positively related to BMI change and %EWL at 6 and 12 months after the surgery [21]. In contrast, a study on 53 patients with at least 6 months post-RYGB suggested carbohydrate intake as an important determinant of weight loss, accounting for 28.5% of the variance in average monthly excess weight loss (AMWL). In that study, a greater AMWL was linked with lower energy (r = -0.373, p = 0.007), carbohydrate (g/day) (r = -0.414, p = 0.003), and fat (g/day) (r = -0.283,p = 0.044) intakes [19].

Among the studies with more than 12 months of followup, one study in 69 patients with a mean time since RYGB of  $30\pm8$  months found that energy and fat intake (% of energy), along with other variables including age, excess weight, and weight before surgery, accounted for 47% of the %EWL [18]. In another study involving 86 patients undergoing SG, energy (r = -0.54) and fat (r = -0.35) were inversely correlated with %EWL at 7 years post-surgery [34]. In contrast, a cross-sectional study in 107 RYGB patients with a mean time since surgery of  $3 \pm 1.8$  years demonstrated that only energy intake (not macronutrients) is associated with %EBMIL [23]. Consistent with the finding, a prospective study on 355 patients undergoing SG and RYGB demonstrated that energy intake, but not the proportion of individual macronutrients, independently predicted %EWL after 5 years [16]. Controlling for relevant confounders, another study showed that a greater decrease in energy intake is associated with a higher %TWL ( $\beta = -0.004$ , p = 0.014) in 135 individuals undergoing SG and RYGB who were followed for more than 4 years [32]. Protein intake 1 year after RYGB was not related to %TWL at the mid-term (2–3 years) or long term (4–5 years) after the surgery in a retrospective study [36].

Two studies defined dietary intake as a categorical variable. One that evaluated the %EWL across three groups of patients with different energy intakes revealed that the mean %EWL of those with a greater energy intake did not differ significantly from those with a lower



Fig. 1 Flowchart of the study selection

energy intake after 1–5 years post-RYGB [22]. Another one compared the mean of %TWL based on protein intake groups of < 60 and  $\geq$  60 g/day that found no significant differences in %TWL between the two groups at 6 and 12 months after SG [29].

## Energy, Macronutrients, and Suboptimal Weight Loss

In four studies, participants were divided based on the rate of weight loss, and then energy and macronutrient intakes

Table 1 Charact	eristics of stud	lies included for v	veight loss after b:	ariatric surgery							
First author	Country	Type of surgery	Study design	Time since surgery	Sample size	Women (%)	Mean age (years)	Outcome	Dietary assess tool	Dietary exposure	Quality <sup>1</sup>
Coughli (1983) [17]	USA	GB	Prospective	1, 3, 6, and 12 months	25	84	32.9	Weight change	Dietary recall	En & macro (g/day & % kcal)	Low
Bobbioni- Harsch (2002) [15]	Switzerland	GB	Prospective	3, 6, and 12 months	50	100	38.4	Weight change	3-day food records	En & macro (% kcal)	Low
Wardé-Kamar (2004) [18]	NSA	RYGB	Prospective	Up to 4 year $(30 \pm 8 \text{ months})$	69	92.8	46	%EWL	24-h dietary recall	En & macro (% kcal)	Low
Faria (2009) [19]	Brazil	RYGB	Retrospective	< 12 months $\& > 12$ months	53	80	36.8	%EWL & AMWL	4-day food record	En & macro (g/day)	Low
Kruseman (2010) [20]	Switzerland	RYGB	Prospective	8±1.2 years	80	100	40	%EWL	4-day food record	En, carbohydrate & fat (%kcal), and protein (g/kg BW)	Low
Ortega (2011) [23]	Spain	RYGB	Cross-sectional	$\geq 1$ years (3.0 $\pm 1.8$ years)	107	79	41.8	%EBMIL	3-day food record	En & macro (g/day)	Low
Raftopoulos (2011) [21]	USA	RYGB	Prospective	3, 6, and 12 months	167	83.3	42.7	%EWL & BMI change	3-day dietary recall	Protein (g/day)	High
Forbush (2011) [22]	USA	RYGB	Retrospective	1–5 years	265	86	48.2	%EWL	FFQ	En	Low
Novais (2012) [24]	Brazil	RYGB	Cross-sectional	$2-7$ years (3.9 $\pm$ 1.4 years)	141	100	44	%EWL	2-day food recall	En & macro (% kcal)	Low
Moizé (2013) [16]	Spain	RYGB &SG	Prospective	Over 5 years	355	75.2	45.4	%EWL	3-day food records & a 24-h dietary recall	En & macro (% kcal)	Low
Sherf-Dagan (2016) [29]	Israel	SG	Prospective	3, 6, 12 months	LL	58.4	42.7	TML%	3-day food diaries	Protein	Low
Amundsen (2017) [30]	Norway	GB	Case-control	> 1 years $(58.3 \pm 23.6 \text{ months})$	49	85.7	46.8	%EWL	FFQ	En & macro (% kcal)	Low
Schoemacher (2019) [32]	Netherlands	RYGB & SG	Prospective	≥4 years (Median=4.8 years)	135	83.7	46.5	TWL	2-day food diary	En & macro (% kcal)	Low
Iossa (2020) [34]	Italy	SG	Prospective	7 years	86	75.6	45	%EWL & %EBMIL	7-day food record	En & fat (g/day)	Low
Holthuijsen (2022) [36]	Netherlands	RYGB	Retrospective	5 year	85	81.2	45.8	TML%	24-h dietary recall	Protein (g/day)	High
<sup>1</sup> Assessed using	the Newcastle-	-Ottawa quality a	ssessment scale								

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AMWL, average monthly weight loss; EBMIL, excess BMI loss; En, energy; EWL, excess weight loss; FFQ, food frequency questionnaire; GB, gastric bypass; Macro, macronutrients; RYGB, Roux-en-Y gastric bypass; SG, sleeve gastrectomy, TWL, total weight loss

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Author, year	Country	Type of surgery	Study design	Time since surgery	WR/non- WR (n)	Women (%)	Mean age, years	WR defini- tion	WR cut-off	Dietary assessment tool	Dietary exposure	Quality <sup>1</sup>
Freire (2012) [25]	Brazil	RYGB	Cross-sec- tional	$45.5 \pm 32.6$ months	56/44	84	45.1	% of weight nadir	≥2%	24-h dietary recall & FFQ	En, macro (%kcal)	Low
Alvarez (2019) [26]	Chile	SG	Case-con- trol	$\geq 2$ years (median = 38.5 months)	20/20	80	42.9	% of maximum weight lost	≥ 25%	FFQ	En, macro (g/day & % kcal)	Low
Reid (2019) [27]	Canada	RYGB	Cross-sec- tional	$\ge 5$ years (12.2 $\pm$ 3.7 years)	17/10	89	53.2	% of total weight loss	≤ 30%	3-day food record	En, macro (g/day & % kcal)	High
da Silva (2019) [28]	Brazil	RYGB	Retrospec- tive	≥ 2 years (47±18 months)	19/61	88.8	46	% of weight nadir	≥ 10%	2-day dietary recall	En & protein (g/day), carbohy- drate & fat (% kcal)	Low
Chou (2017) [31]	Taiwan	SG	Retrospec- tive	≥5 years	9/28	75	33.5	% of weight nadir	≥ 25%	DQES v2	En, macro (g/day & % kcal)	Low
Amundse (2017) [30]	Norway	GB	Case-con- trol	<ul> <li>&gt; 1 years</li> <li>(58.3 ± 23.7 months)</li> </ul>	36/11	89.4	46.9	% of maximum weight lost	> 15%	FFQ	En, macro (% kcal)	Low
Vieira (2018) [ <b>33</b> ]	Brazil	RYGB	Cross-sec- tional	> 2 years (67.1 ± 33.1 months)	20/20	100	40.1	% of weight nadir	≥ 10%	24-h dietary recall and 2-day food record	En, macro (g/day)	Low
Iossa (2020) [34]	Italy	SG	Retrospec- tive	7 years	24/48	88.9	45	% of EWL	≥25%	7-day food record	En, macro (g/day)	Low
Damin (2021) [ <b>35</b> ]	Brazil	RYGB	Cross-sec- tional	≥5 years (6.3±0.81 years)	22/51	100	42.6	% of maximum weight lost	≥ 10%	3-day food record	En, macro (% of energy)	High
Jabbour (2021) [37]	Lebanon	RYGB & SG	Cross-sec- tional	≥6 months (2.4±1.8 years)	24/36	85	35.5	Any weight regain from nadir weight	None	2-day food record	En	Low

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were compared across the groups of participants [18, 20, 24, 30]. Participants in the four studies underwent RYGB. All of the studies defined SWL as 50% of EWL and looked at macronutrients as a percentage of energy, except for one, which used g/kg body weight to measure protein intake [20]. Patients were divided into two groups (< 50% and  $\ge 50\%$  of EWL) in three studies [18, 20, 30] and three groups (<50%, 50–75%, and  $\geq$  75% of EWL) in one study [24]. In one study, although the energy intake of the SWL group was higher than that of the AWL group at 18–47 months post-surgery, the differences did not reach significance [18]. The other study found that women with SWL had higher energy intake at 8 years postoperative than those with AWL  $(1934 \pm 501)$ vs.  $1634 \pm 526$  kcal/day, p = 0.02), but energy intakes at 1 year postoperative did not differ significantly [20]. There were no significant differences in energy intake between the two groups of patients with SWL and AWL at > 1 year or 2–7 years post-surgery [24, 30]. None of the studies found significant differences in macronutrient contributions to energy intake across the groups of EWL. For the metaanalysis, we pooled the data of the two groups of 50–75% and > 75% EWL in the Novais et al. study [24] to generate a group for individuals with AWL consistent with the other studies. As it has been shown in Fig. 2, a meta-analysis of the 4 studies (96 SWL/227 AWL) indicated a significantly higher energy intake in SWL patients than in AWL patients (MD = 202.5 kcal/day, 95% CI = 77.3, 327.7;  $I^2 = 0\%$ , p heterogeneity = 0.691). The Egger test showed no evidence for small-study effects (p = 0.774). Meta-analyses of the studies revealed no significant differences in macronutrient composition as a percentage of energy between the two groups (Table 3).

#### **Energy Intake and Weight Regain**

Ten studies compared daily energy intakes between the WRs and non-WRs [25–28, 30, 31, 33–35, 37]; nine of them found that WRs consumed more energy [25–28, 30, 33–35,



Fig. 2 Forest plot of mean differences in energy intake between suboptimal weight loss and acceptable weight loss groups

Table 3 Pooled mean differences in macronutrient intakes between patients with postoperative suboptimal weight loss and acceptable weight loss

Macronutrients intakes	No. study	No. of participants <sup>1</sup>	MD (95% CI)	<i>I</i> <sup>2</sup> (%)	<i>p</i> heterogeneity	Egger's test
Carbohydrate (% of energy)	3	76/192	-0.76 (-2.92, 1.41)	11	0.325	0.873
Fat (% of energy)	3	76/192	1.76 (-0.51, 4.04)	36.1	0.209	0.362
Protein (% of energy)	2	43/145	-0.76 (-1.98, 0.46)	0	0.882	-

<sup>1</sup>Suboptimal/acceptable weight loss groups. CI, confidence interval; MD, mean difference

37]. Two studies' findings were statistically significant [25, 34], six were not [26–28, 30, 33, 35], and one did not provide a *p* value for the comparison [37]. Contrary to the other research, one found that WRs had a non-significant lower energy intake than non-WRs [31]. The forest plot for MD in energy intake between WRs and non-WRs is illustrated in Fig. 3. The meta-analysis of the 10 studies (238 WRs/323 non-WRs) showed a significantly higher intake of energy in WRs (MD = 192 kcal/day, 95%CI=95.6, 288;  $I^2$  = 37.8%, *p* heterogeneity = 0.106). There was no publication bias by Egger's test (*p*=0.716). Subgroup analysis based on surgery type showed the difference was non-significant for SG, but there was high heterogeneity =0.011).

#### **Carbohydrate and Weight Regain**

Nine studies compared daily carbohydrate intake between WRs and non-WRs [25–28, 30, 31, 33–35]. Four of them look at carbohydrates as a percentage of energy intake [25, 28, 30, 35], two as absolute intake (g/day) [33, 34], and three as both [26, 27, 31]. One of the seven studies examining

carbohydrates as a percentage of energy found that WRs had a significantly higher carbohydrate intake than non-WRs (57.1% vs. 48.2%; p value < 0.001) [35], while the other studies showed no significant differences. Furthermore, in one of the five studies comparing carbohydrate consumption in g/day, WRs ingested more carbohydrates than non-WRs  $(222 \pm 84.3 \text{ vs. } 162 \pm 67.5 \text{ g/d}; p \le 0.05)$  [27]. Meta-analysis of seven studies indicated that the proportion of carbohydrates to energy intake did not significantly differ between WRs and non-WRs. However, the absolute intake of carbohydrates was 24.9 g/day (95%CI = 5.98, 43.8;  $I^2$  = 64.4%, p heterogeneity = 0.024) more in WRs vs. non-WRs according to the pooled analysis of five studies. Carbohydrate intake was also significantly greater in WRs undergoing SG, according to a subgroup meta-analysis by surgery type (Table 4).

#### **Fat and Weight Regain**

Of the nine studies that compared fat consumption in the two groups of WRs and non-WRs, four looked at fat as a percentage of energy [25, 28, 30, 35], two looked at fat in



Fig. 3 Forest plot of mean differences in energy intake between weight regain and non-weight regain groups

 Table 4
 Pooled mean differences in macronutrient intakes between weight regain and non-weight regain groups

Dietary intake	No. study	No. participants <sup>1</sup>	MD (95% CI)	$I^{2}(\%)$	p heterogeneity	Egger's test
Carbohydrate(% of energy	y)					
All	7	179/225	1.53 (-2.39, 5.48)	84.6	< 0.001	0.723
Gastric bypass	5	150/177	1.44 (-3.70, 6.58)	87.1	< 0.001	0.899
Sleeve gastrectomy	2	29/48	1.70 (-6.43, 9.83)	86.9	0.006	-
Carbohydrate(g/day)						
All	5	81/120	24.9 (5.98, 43.8)	64.4	0.024	0.411
Gastric bypass	2	28/24	27.0 (-21.6, 75.5)	62.4	0.103	-
Sleeve gastrectomy	3	53/96	26.3 (2.40, 50.2)	61.1	0.077	0.578
Fat(% of energy)						
All	7	179/225	-0.17 (-3.12, 2.77)	78.9	< 0.001	0.474
Gastric bypass	5	150/177	-0.67 (-4.44, 3.11)	82.1	< 0.001	0.723
Sleeve gastrectomy	2	29/48	1.04 (-5.03, 7.11)	80.6	0.023	-
Fat(g/day)						
All	5	81/120	4.52 (-6.77, 15.8)	75.6	0.003	0.669
Gastric bypass	2	28/24	1.99 (-8.78, 12.8)	0	0.667	-
Sleeve gastrectomy	3	53/96	5.36 (-12.6, 23.4)	86.6	< 0.001	0.827
Protein(% of energy)						
All	6	160/164	-1.46(-2.52, -0.41)	9	0.359	0.390
Gastric bypass	4	131/116	-1.27 (-2.52, -0.02)	11.6	0.335	0.538
Sleeve gastrectomy	2	29/48	-2.02 (-4.57, 0.053)	39.1	0.200	-
Protein(g/day)						
All	6	100/181	-3.59 (-11.2, 4.01)	35.3	0.172	0.584
Gastric bypass	3	47/85	-2.73 (-13.0, 7.50)	0	0.943	0.145
Sleeve gastrectomy	3	53/96	-5.48 (-21.4, 10.4)	73.7	0.022	0.710

<sup>1</sup>Weight regain/non-weight regain groups. CI, confidence interval; MD, mean difference

g/day [33, 34], and three looked at both [26, 27, 31]. The percent of energy from fat was not significantly different between WRs and non-WRs in four studies [25, 27, 28, 30], but in two studies, the percent was significantly lower in WRs [31, 35]. Considering absolute fat intake as g/day, Chou et al. study also found that WRs consumed less fat per day than non-WRs  $(40 \pm 12.8 \text{ vs. } 55 \pm 27 \text{ g/day}; p = 0.03)$ [31]. In contrast, studies by Alvarez et al. and Iossa et al. found that WRs consumed more fat [26, 34]. Two studies did not demonstrate any significant difference in the daily intake of fat as g/day [27, 33]. The pooled effect size of seven studies (179 WRs/225 non-WRs) for percent of energy from fat and five studies (81 WRs/120 WRs) for fat as g/day represented no significant differences between the groups. In addition, no significant differences in fat intake were found in the subgroup analysis (Table 4).

#### **Protein and Weight Regain**

Daily protein intake in three studies was reported as a percentage of daily energy intake [25, 30, 35], in three studies reported as g/day [28, 34], and in three studies reported as both [26, 27, 31]. The seven studies could not show any significant difference in protein intake between the groups, either as a percent of energy from protein or g/day [25–28, 30, 33, 34]. Chou et al. found that WRs consume less protein than non-WRs ( $52.9 \pm 17.5$  vs.  $77.5 \pm 35.3$  g/d; p = 0.01), but the difference was not significant when considered as percent energy from protein  $(20.3 \pm 3 \text{ vs. } 23.6 \pm 5.3\%)$  of energy; p = 0.08 [31]. Moreover, WRs compared to non-WRs had a higher percentage of energy from the protein in the Damin et al. study (median (interquartile range), 15.2 (13.2-18.9) vs. 17.4 (14.5–22.2); p = 0.054) [35]. The pooled effect size of six studies (Table 4) demonstrated that the proportion of protein in daily energy intake was slightly lower in WRs than in non-WRs (MD = -1.46%, 95% CI = -2.52, -0.41;  $I^2 = 9\%$ , p heterogeneity = 0.359), but there was no significant difference in protein intake as g (MD = -3.59, 95%CI = -11.2, 4.01;  $I^2 = 35.3\%$ , p heterogeneity = 0.172).

## Discussion

Previous studies on the associations between postoperative intakes of energy and macronutrients with weight loss have employed three distinct approaches: considering both dietary intake and weight loss as continuous variables [15-19, 21, 23, 32, 34, 36]; comparing weight loss at different levels of dietary intake as a categorical variable [22, 29]; and comparing food intake at different levels of weight loss as a categorical variable [18, 20, 24, 30]. Studies using the first approach consistently showed that higher energy intake was associated with lower weight loss in both the short and long term after surgery. However, the results of the investigations into the relationship between macronutrients and weight loss were inconclusive. Some studies found no association between macronutrients and weight loss [15, 16, 23, 32], while others found that fat [17, 18, 34] or carbohydrates [19] were inversely associated with weight loss. In addition, one of the two studies on protein intake (g/day) and weight loss reported positive associations between protein intake and %EWL at 6 and 12 months post-RYGB [21], while the other found no significant associations between protein intake at 1 year postoperative and %TWL at mid-term and long term after RYGB [36]. We were unable to do meta-analyses since the studies were so diverse.

Studies employing the second approach demonstrated no significant differences in %EWL between the three groups of participants with varying energy intake 1–5 years after RYGB [22] or in %TWL between those with a protein intake of  $\geq$  60 g/day and those with a protein intake of < 60 g/day over 12 months after SG [29].

The results of the four investigations comparing the energy intake of patients with SWL and AWL using the third approach were likewise inconsistent [18, 20, 24, 30]. However, none of them could find significant differences in macronutrient intakes between SWL and AWL. According to the meta-analysis done in this study, SWL consumed 203 kcal/day (95% CI = 77, 328) more energy than AWL, although there were no differences in macronutrient consumption between the two groups.

The majority of prior research reported a greater energy intake in WRs than in non-WRs, but the difference was statistically significant only in two studies [25, 34]. Our meta-analysis indicated that WRs consume more energy (MD = 192 kcal/day; 95% CI=95, 288) and carbohydrates (MD=25 g/day; 95% CI=5.98, 43.8) than non-WRs. We also found that the proportion of protein to energy in WRs is slightly lower than in non-WRs (MD = -1.46, 95% CI=-2.52, -0.41).

In some studies, the postoperative time of participants included in a study was widely different [16, 19, 22–25, 30, 33] which may be a source of bias in their findings because the time since surgery is a predictor of weight loss after surgery [11, 16]. In addition, dietary intake may also change over time after surgery. If there was a temporal variation in both weight outcomes after surgery and energy and macro-nutrient intakes, future studies on the association of energy

and macronutrient intakes with weight outcomes should be conducted in a more homogenous population.

A recent systematic review suggested 11 risk factors from five main categories for weight regain. Increased sweet consumption and portion size as well as eating behaviors were dietary risk factors for weight regain [11]. The systematic review also included 5 studies published until July 2019 on the association between energy and macronutrients and weight regain, but due to the mixed results of the studies, no conclusion was made. The systematic review only included studies with a predefined threshold of  $\geq 10\%$  for weight regain. Although the definition of WR has been different across the studies, we decided not to consider any exclusion criteria according to the definition for the following two main reasons. First, the standard definition and threshold for weight regain after BS have not been established yet. Second, findings of a prospective study showed that all the different weight regain measures after RYGB are associated with an increased risk of diabetes progression, declines in physical health-related quality of life, and satisfaction with surgery [38]. Indeed, the results of that study imply that any weight regained after BS may be clinically important and should not be neglected. Therefore, in our systematic review, two more studies published before July 2019 and 3 more studies that were published after the date were included compared to the earlier systematic review. Furthermore, we performed meta-analyses to make a clear conclusion on the association between energy and macronutrient intake after surgery and WR, which has not been done previously. To our knowledge, no systematic review has looked at how energy and macronutrient intakes are related to weight loss after BS.

A strength of the current study is that it covers both SWL and weight regain, the two distinct types of weight loss failure after BS [9]. Therefore, this study provides a comprehensive overview of the association between energy and macronutrient intakes and post-surgery weight outcomes after SG and GB. We also conducted meta-analyses to determine how much energy and macronutrients in individuals with SWL or WR consumed differently compared to individuals with AWL or non-WR.

When interpreting the findings of this study, its limitations should be kept in mind. First, only a few studies were included for the comparison of food intakes between SWL and AWL, and the surgery types were all GB. Second, the number of studies included to study weight regain was limited, and most of them were restricted by sample size and therefore the power to detect an association. Third, weight regain is defined as gaining weight following initial successful weight loss (EWL > 50%) [9]. However, WRs in some studies had a mean %EWL of less than 50% at the time of the study [27, 34]. Therefore, it is unclear if they achieved a satisfactory %EWL following surgery. Fourth, since the dietary assessment tools are self-reported, the possibility of under-reporting, particularly in those regaining weight, is high. An additional limitation was that, due to a paucity of data, we were unable to do subgroup meta-analyses based on the time after surgery and the weight regain definition. In addition, subgroup meta-analyses based on surgery types for weight regain revealed that the general directions of associations for SG and GB were comparable, but the significant findings in MD of energy, carbohydrate (g), and protein (% of energy) did not reach significance in SG for energy and percentage of protein and GB for absolute intake of carbohydrate. The lack of significance in one of the categories may be attributable to the limited number of studies, their small sample sizes, and the variability of the results among the studies.

In conclusion, GB patients with SWL consumed 203 kcal/ day more energy than those with AWL, but the proportion of macronutrients to energy intake did not differ significantly between the two groups. In addition, WRs consumed an average of 192 kcal/day more energy and 25 g/day more carbohydrates than non-WRs, but the percentage of protein intake was 1.46% lower. Since there is no strong evidence supporting the association of dietary macronutrient compositions and weight outcomes after BS, it seems that energy intake, independent of macronutrient proportions, is crucial for weight reduction and prevention of weight rebound after GB or SG. Further studies with a larger sample size and a more homogenous population are needed to clarify the role of energy and macronutrients on weight outcomes after BS.

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**Data Availability** Data that support the findings of this study are available from the corresponding authors upon reasonable request.

#### Declarations

**Ethics Approval** This article does not contain any studies with human participants or animals performed by any of the authors.

Consent to Participate Informed consent does not apply.

Conflict of Interest The authors declare no competing interests.

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