## **RESEARCH**





# **Metabolic‑Associated Fatty Liver Disease and Weight Loss After Bariatric Surgery: A Systematic Review and Meta‑Analysis**

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## **Abstract**

**Background** Metabolic Dysfunction-Associated Steatotic Liver Disease (MASLD) and Metabolic Dysfunction-Associated Steatohepatitis (MASH) are increasingly prevalent in patients undergoing bariatric surgery (BS). Understanding their impact on weight loss outcomes after surgery and highlighting the results of surgical techniques such as Roux-en-Y Gastric Bypass (RYGB) and Sleeve Gastrectomy (SG) in relation to the presence of MASH are essential for improving patient management and predicting long-term success.

**Methods** A systematic review and meta-analysis were conducted. We searched the PubMed database; inclusion criteria were BS patients with liver impairment data at surgery and weight loss data at follow-up of 6 months or longer. Meta-analyses were conducted using R's meta package, assessing heterogeneity with the  $I^2$  statistic and employing subgroup analyses where necessary.

**Results** Out of 1126 eligible studies, 22 were included in the fnal systematic review. For the MASLD vs. Normal Liver (NL) comparison, no signifcant diference in BMI change was found at 12 months, but subgroup analysis indicated a possible publication bias (published data vs data collected). In the MASH vs. non-MASH comparison, high heterogeneity was noted at 12 months, and further stratifcation by surgical technique revealed that SG patients with MASH experienced lower weight loss, approaching statistical signifcance.

**Conclusions** MASLD does not signifcantly afect short-term weight loss outcomes post-BS, but long-term results show variability. Standardized reporting practices and complete data dissemination are essential for future research to enhance meta-analysis reliability and generalizability.

**Key points**

- MASLD does not signifcantly impact short-term weight loss post-bariatric surgery.
- MASH patients with sleeve gastrectomy may lose less weight in the frst year.
- High study heterogeneity highlights the need for standardized reporting.
- Long-term outcomes vary, stressing the need for preoperative liver assessment.

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### **Graphical Abstract**



Metabolic-associated fatty liver disease and weight loss after bariatric surgery:

**Keywords** Weight loss · NAFLD/MASLD · MASH · Bariatric surgery · Sleeve gastrectomy · Roux-en-Y gastric bypass

# **Introduction**

Obesity is a chronic disease defned by a body mass index (BMI) greater than 30 kg/m<sup>2</sup>, and its prevalence is increasing exponentially worldwide, creating a signifcant global health problem [\[1](#page-10-0)]. Classifed as a pandemic, it leads to the development of type 2 diabetes mellitus (T2DM), metabolic syndrome, and fatty liver disease, the latter recently renamed as MASLD (Metabolic dysfunction-Associated Steatotic Liver Disease), among other associated problems [[2,](#page-10-1) [3\]](#page-10-2). MASLD represents a clinical syndrome caused by excessive fat deposition in hepatocytes and includes histopathological entities ranging from simple steatosis (SS) and metabolic dysfunction associated steatohepatitis (MASH) to its advanced stages, including hepatic fbrosis and cirrhosis [[4](#page-10-3)]. It is a highly prevalent disease afecting approximately 25% of the global population but can increase up to 60% in diabetic patients and up to 90% in individuals with severe obesity [\[5](#page-10-4)]. Bariatric surgery (BS), in addition to inducing signifcant weight loss in patients with obesity, has also been associated with histological improvement of SS, MASH, and even partial regression of fbrosis in early cases [\[6](#page-10-5)]. Furthermore, BS is also associated with a substantial reduction in the risk of progression from MASH to cirrhosis (88% according to recently published data [\[7](#page-10-6)]). However, few studies have analyzed the relationship between weight loss after BS based on the presence and stage of MASLD. A recent study highlights those patients without underlying liver disease lose more weight than those with low- or high-grade liver disease after sleeve gastrectomy (SG) [\[8](#page-10-7)]. There have been slight diferences in the percentage of excess weight loss (%EWL) 12 months after gastric bypass (RYGB) between patients with MASH and those with SS. While both means exceeded 50% of their value, this does not discriminate

between possible diferences that may exist due to the bias exerted by the initial BMI value [\[9\]](#page-10-8). Weight regain after surgery can worsen MASLD or delay its improvement, and clinically, it is also important to refne the indication of the appropriate surgical technique based on the patient's liver histology. In this sense, patients with MASLD/MASH could beneft more from techniques that achieve a sustained weight loss pattern, as seen with RYGB [\[10](#page-10-9), [11](#page-11-0)].

Regarding diferent indicators of weight loss after BS, %EWL is more sensitive to error than total weight loss  $(\%TWL)$  and adjustable weight loss (%AWL), as it is closely related to the ideal weight. %TWL is a parameter that should be used in all scientifc communications and publications, and although we do not yet have sufficient evidence regarding %AWL, it seems to be even more precise than %TWL [[12](#page-11-1)]. %TWL allows for comparisons between different patient series with less bias from initial BMI, and %AWL is also very useful in cohorts of diabetic patients [\[13,](#page-11-2) [14](#page-11-3)]. A recent study demonstrates that after analyzing weight loss through these indicators, both diabetic patients undergoing RYGB and those undergoing SG showed no diferences in weight loss in the presence of MASH. On the other hand, non-diabetic patients undergoing SG lost less weight in the presence of MASH compared to patients without MASH; however, the presence of MASH did not affect weight loss in non-diabetic patients undergoing RYGB. Therefore, there seem to be relevant diferences in weight loss patterns between patients with or without MASH, especially after SG [[15\]](#page-11-4).

The objective of this meta-analysis is to analyze the effect of the presence of MASLD and MASH on weight loss after BS using the two most frequently used techniques worldwide (SG and RYGB) and based on diferent weight loss indicators.

# **Materials And Methods**

## **Data Sources**

For the purpose of conducting this systematic review and meta-analysis, our main research question was formulated as "Is weight loss after bariatric surgery infuenced by the presence of MASLD/ MASH?" Table [1](#page-2-0) depicts the population, intervention, comparator, and outcomes (PICO) structure of information. The systematic review was registered at PROS-PERO [\(https://www.crd.york.ac.uk/prospero/](https://www.crd.york.ac.uk/prospero/)).

To fully capture the literature related to this topic, we performed two rounds of searches in the PubMed [\(https://pub](https://pubmed.ncbi.nlm.nih.gov/)[med.ncbi.nlm.nih.gov/](https://pubmed.ncbi.nlm.nih.gov/)) online database, the frst one used the query "((bariatric surgery[Title/Abstract]) OR (Sleeve Gastrectomy[Title/Abstract]) OR (gastric bypass[Title/ Abstract])) AND ((liver[Title/Abstract]) OR (nafd[Title/ Abstract]) OR (MASH[Title/Abstract]) OR (masld[Title/ Abstract]) OR (mash[Title/Abstract])) AND (weight[Title/ Abstract])" and included studies from 2018 onwards. The second search used the query "("bariatric surgery" OR "Sleeve Gastrectomy" OR "gastric bypass" OR "gastric band") AND ("fatty liver" OR nafd OR MASH OR masld OR mash) AND (weight OR BMI OR EWL OR TWL)," without any date specifcations. Both searches included the human and adult flters. In addition, we also inspected other sources, such as references of relevant publications. To avoid publication bias, we intentionally included studies whose main outcome was not the assessment of weight loss after bariatric surgery but reported this data incidentally.

## **Study Selection**

Study inclusion criteria were bariatric surgery patients, the existence of two groups based on liver impairment, and weight loss data at any follow-up 6 months or longer after the surgery. Exclusion criteria included the lack of follow-up weight loss data, case reports, reviews or guidelines, animal studies, and fbrosis and cirrhosis comparisons. Given the vast majority of studies that did not provide weight loss data stratifed by initial liver status, we opted to contact the authors requesting it. We did not request any data from

studies prior to 2014, due to the low likelihood of getting answers.

We did not exclude any papers based on language but rather used the online tool DocTranslator ([www.onlinedoct](http://www.onlinedoctranslator.com) [ranslator.com\)](http://www.onlinedoctranslator.com) to assess their eligibility.

The quality of the studies was evaluated independently by two authors (V.A. and M.V.) using the scale described by Qumseya [[16](#page-11-5)].

## **Data Synthesis**

Several studies presented data as medians and quartile ranges. We used the quantile estimation method for estimating the mean and standard deviation using the R  $(v3.6.1)$ package estmeansd (v 1.0.1) [[17\]](#page-11-6). For several reports, means and standard deviations had to be combined (for instance, simple steatosis and MASH groups both combined into a MASLD group, which encompassed the broad spectrum of the disease), using the following formulas, where *m* is the mean (c for combined, 1 and 2 for the groups to combine),  $\sigma$ is the standard deviation, and  $n_1$  and  $n_2$  are the samples sizes for both groups:

$$
m_c = \frac{n_1 \cdot m_1 + n_2 \cdot m_2}{n_1 + n_2} \quad \sigma_c = \sqrt{\frac{(n_1 - 1) \cdot \sigma_2^1 + n_1 \cdot m_1^2 + (n_2 - 1) \cdot \sigma_2^2 + n_2 \cdot m_2^2 - (n_1 + n_2) \cdot m_c^2}{n_1 + n_2 - 1}}
$$

Finally, given that some studies indicated BMI at baseline and at follow-up, but did not include directly the ∆BMI with the corresponding standard deviation, we had to infer the standard deviation. For this, we used the formula described by McNemar [[18](#page-11-7)] with a correlation of 0.6 (estimated from Salman et al.'s [[19\]](#page-11-8) raw data).

#### **Meta‑Analysis**

Studies were grouped according to liver impairment histological classifcation ([[1\]](#page-10-0) normal liver (NL) group versus MASLD and [\[2](#page-10-1)] non-MASH versus MASH), weight measurement, and follow-up time. When follow-up time was not equal for all study participants, the study was assigned to the closest time mark.

We used R's package meta (v7.0–0, [[20\]](#page-11-9)), which calculates common and random efect estimates using inverse variance weighting for pooling. Efect sizes were assessed with

<span id="page-2-0"></span>**Table 1** Researchable question defnition via the PICO structure

Population	Human subjects of any age and any degree of obesity (but having obesity) undergoing bariatric surgery and with liver diagnosis
Intervention	Bariatric surgery, later limited to RYGB and SG
Comparison	Patients with and without MASLD (diagnosed by either liver biopsy, imaging or ultrasound techniques) and patients with and without MASH (diagnosed by liver biopsy)
<b>Outcomes</b>	Weight loss measurements after bariatric surgery; change in BMI, TWL, EWL, and weight at different time points

forest plots, and study heterogeneity was evaluated with the  $I<sup>2</sup>$  statistic and funnel plots. The random effects model was used when  $I^2$  > 0.05; otherwise, the common effect model was applied. Sub-group analyses were performed in order to find the source of heterogeneity in instances when  $I^2 > 0.05$ . *P*-values < 0.05 were considered statistically significant.

Variables used for sub-group analyses included data source (original report or email), surgery type, a statistically signifcant diference in age or initial BMI between groups, or more than 15% diference in sex distribution between groups.

# **Results**

## **Identifed Records**

We retrieved a total of 1126 eligible studies; 1032 records were screened and 247 were assessed for eligibility. From these, sixteen studies provided the data in the paper or supplementary materials, while 140 indicated that the data was available but did not include it in the original paper (generally, weight loss data was not stratifed by liver status at baseline). Out of the 108 emails sent to the authors, responses were received for 13, seven of them provided the data, and fnally, six studies were included (one study [[21\]](#page-11-10) was discarded due to having classifed patients only according to NAFLD Fibrosis Score, NFS).

As depicted in Fig. [1A](#page-4-0), a total of 22 studies were included in the fnal meta-analysis. We assessed the possible study combinations regarding liver histological classifcation, weight measurement, and time points. For the MASLD vs.  $NL$  comparison (Fig. [1](#page-4-0)B), sufficient records were found for the change in BMI at 6 and 12 months, as well as for EWL and TWL at 12 months. For the MASH vs. non-MASH comparison (Fig. [1C](#page-4-0)), the change in BMI can be assessed at 6, 12, and 24 months, and EWL and TWL can be evaluated at 12 months. No other weight loss measurements were widely available in the screened reports. Table [2](#page-5-0) reports all specifcations from the included studies.

## **MASLD Impact on Weight Loss After Bariatric Surgery**

First, we aimed to evaluate the impact of MASLD presence on weight loss after bariatric surgery. As described in Fig. [1b](#page-4-0), we performed a meta-analysis for BMI change at 6 and 12 months, and for EWL and TWL at 12 months. The four studies reporting a change in BMI at 6 months had low heterogeneity  $(I^2 = 0\%)$  and showed no statistically significant diference in weight loss patterns between patients with MASLD and with NL. Regarding the change in BMI 1 year after surgery, the seven studies were heterogeneous (Fig. [2a](#page-7-0)); the sub-group analysis based on data source showed that there was a statistically signifcant diference only in the studies that directly reported this data in the raw paper (Fig. [2](#page-7-0)b), but this could not be found in the papers that did not directly report this data (Fig. [2](#page-7-0)c). Regarding EWL and TWL at 12 months, we did not fnd any diference.

## **MASH Impact on Weight Loss After Bariatric Surgery**

On the one hand, studies reporting changes in BMI at 6 months had a low heterogeneity, and we did not fnd any impact of MASH presence on BMI loss at this time point. On the other hand, studies reporting a change in BMI at 1 year initially showed no diference and exhibited high heterogeneity (Fig. [3](#page-8-0)a). Specifcally, the study by Anjani et al. [[34\]](#page-11-11) markedly differs from others. Given that the MASH patients from this study had signifcantly diferent initial BMI compared to the non-MASH patients, we excluded studies with this circumstance. The remaining ten studies displayed high heterogeneity (Fig. [3](#page-8-0)b), and the data source did not explain the cause of this heterogeneity. Therefore, we opted to perform sub-group analyses separately for the two main bariatric surgeries: RYGB (Fig. [3c](#page-8-0)) and SG (Fig. [3](#page-8-0)d). In this case, a diferent trend is observed with the SG group's studies, approaching statistical signifcance in the context of MASH. At 1 year, EWL and TWL were not statistically diferent between MASH and non-MASH patients.

Finally, we assessed the change in BMI 2 years after the bariatric surgery for all studies reporting this data (Fig. [4a](#page-9-0)). The heterogeneity in these studies was very high, and the funnel plot (Fig. [4b](#page-9-0)) showed that the study by Chisholm et al. [[36\]](#page-11-12) differed from the others, being the only one including patients who undergone LAGB. Therefore, we chose to remove it and assess only studies including SG and RYGB patients, obtaining no diference in BMI change at 24 months after bariatric surgery (Fig. [4](#page-9-0)c) between patients with MASH and those without the condition. We could not perform sub-group analysis based on surgery type, as there were not enough studies reporting data separately.

# **Discussion**

Individual studies on weight loss after bariatric surgery in patients with MASLD may yield inconsistent or contradictory results due to diferences in the types of bariatric surgeries, the severity of MASLD, and the patient's characteristics (age, gender, comorbidities, etc.). The results could have important implications for clinical decision-making, as if patients with severe hepatic impairment tend to lose less weight, it may be necessary to adapt pre- and postoperative strategies to optimize their outcomes or refne the indication for the main surgical techniques. It can also help to <span id="page-4-0"></span>**Fig. 1** Study selection and grouping. **a** PRISMA diagram for the studies included in the systematic review, with exclusion reasons. **b** Studies included in the MASLD vs. NL comparison; **c** studies included in MASH vs. non-MASH comparison. Boldface highlighted cells mark the weight measurement and time points analyzed



inform patients about what to expect in terms of long-term outcomes. This has been the hypothesis that has guided the need to conduct a meta-analysis on the subject.

Focusing on the discussion of the results, the absence of signifcant diferences in BMI changes at 6 months suggests that MASLD does not impact early post-surgical weight loss. However, the high heterogeneity and variability in BMI changes observed at 12 months indicate that longterm outcomes could be afected by various factors related to the methodologies used in the analyzed studies, including study design, data reporting, and patient characteristics. That is why we contacted numerous authors, and from the data received, it was observed that there is a clear publication bias in this regard. The manner in which data is reported can substantially affect the perceived efficacy of bariatric surgery in terms of changes in BMI. Specifcally, variations in reporting practices—including the statistical measures used (e.g., mean versus median BMI changes), follow-up durations, sample characteristics, defnitions of success, and publication biases—can all infuence the interpretation of surgical outcomes. The same issues are observed in other studies that analyze the impact of revisional bariatric surgery on weight loss [[39\]](#page-11-13). Examining exclusively the published data for patients with and without MASLD, we observe that individuals with preexisting MASLD experienced signifcantly less total and excess weight loss compared to those

<span id="page-5-0"></span>





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(c)		<b>Experimental</b>			Control				<b>Standardised Mean</b>			Weight	Weight			
	<b>Study</b>	<b>Total Mean</b>			<b>SD</b> Total Mean		<b>SD</b>		<b>Difference</b>			<b>SMD</b>		95%-CI (common) (random)		
	Khalaj et al (2020)			668 15.60 4.3000			132 15.50 4.7000						$0.02$ [-0.16; 0.21]	50.4%	50.4%	
	Blume et al (2021)			308 16.37 5.8500			153 16.29 4.9100						$0.01$ [-0.18; 0.21]	46.7%	46.7%	
	Giraudi et al (2021)			64 14.02 5.4100			7 12.80 5.3600						$0.22$ [-0.56; 1.00]	2.9%	2.9%	
	Common effect model 1040				292								$0.02$ [-0.11; 0.16]	100.0%		
	<b>Random effects model</b>												$0.02$ [-0.08; 0.13]		100.0%	
	<b>Prediction interval</b> Heterogeneity: $I^2 = 0\%$ , $\tau^2 = 0$ , $p = 0.88$											$[-0.83; 0.88]$				
	Test for overall effect (common effect): $z = 0.36$ ( $p = 0.72$ )							$-1$	$-0.5$		0.5					

<span id="page-7-0"></span>**Fig. 2** Meta-analysis for the diference in BMI between baseline and 1 year after bariatric surgery for MASLD (experimental group) vs. NL (control group). **a** Combined results and **b** papers describing this

without MASLD. This diferential outcome was consistent across both surgical techniques employed (RYGB and SG) [\[8,](#page-10-7) [23](#page-11-15)]. The impact of MASH on weight loss after bariatric surgery was also analyzed, focusing on BMI changes at diferent time points. There was no signifcant impact of MASH on the amount of BMI lost at 6 months. The studies reporting BMI changes at 6 months exhibited low heterogeneity, indicating that the results were relatively consistent across these studies. This suggests that, in the short term, the presence of MASH does not afect the weight loss outcomes following bariatric surgery. However, studies reporting BMI changes at 1 year showed high heterogeneity. Despite excluding studies with signifcant initial BMI data in the publications or supplementary materials; **c** authors provided data a *posteriori*

diferences between groups, the remaining ten studies still displayed high heterogeneity. Despite these results, one of the studies that could not be included in the meta-analysis due to the available follow-up time concluded that patients with MASH showed less weight loss starting at 24 months after surgery [\[22\]](#page-11-14). However, another study found no signifcant diferences in weight loss between the MASH and non-MASH groups at a 10-year follow-up [[28\]](#page-11-20). This ongoing variability suggests that other factors, such as the type of surgical technique performed, may infuence the results. In this context, when stratifying the studies according to the surgical technique, it is observed that, in the case of SG, patients with MASH experienced a lower degree of



**Standardised Mean Difference** 







<span id="page-8-0"></span>**Fig. 3** Meta-analysis for the diference in BMI between baseline and 1 year after bariatric surgery for MASH (experimental group) vs. non-MASH (control group). **a** Combined results, including studies where initial BMI is signifcantly diferent between groups; **b** metaanalysis of studies with no signifcantly diferent initial BMI; **c** metaanalysis of only RYGB studies; and **d** meta-analysis of only SG studies

weight loss very close to statistical signifcance, compared to those without MASH. This is consistent with the results found in our recently published patient series [[15](#page-11-4)]. But histopathologically, no signifcant diferences have been observed between SG and RYGB after surgery in a published meta-analysis with controlled studies, which seems



 $-1.5$ 

 $-1$   $-0.5$  0

 $0.5 \quad 1$ 

 $1.5$ 

**Prediction interval** Heterogeneity:  $l^2 = 76\%$ ,  $\tau^2 = 0.0996$ ,  $p < 0.01$ 

Test for overall effect (common effect):  $z = -2.46$  ( $p = 0.01$ ) Test for overall effect (random effects):  $t_3 = -0.16$  ( $p = 0.89$ )

<span id="page-9-0"></span>**Fig. 4** Meta-analysis for the diference in BMI between baseline and 2 years after bariatric surgery for MASH (Experimental group) vs. non-MASH (Control group). **a** Combined results; **b** funnel plot show-

to indicate that both procedures may be equally efective in the management of MASH  $[11]$  $[11]$ . This raises the question of whether the diference in weight loss can largely be attributed to the surgical technique used. Undoubtedly, having more data would help us confrm or refute this hypothesis SG, also seems to be safer than other procedures in patients with cirrhosis [[40\]](#page-11-29). This can contribute to an increase in the number of SG performed as shown in the IFSO surveys [\[41\]](#page-11-30), and because of this, the weight results could be not so optimal. We do not know what will happen to liver function in the long term. Recent studies have determined that bariatric surgery reduces the risk of non-alcoholic cirrhosis and liver cancer but may also increase the risk of postoperative ing study heterogeneity; and **c** meta-analysis of studies with patients undergoing either SG or RYGB

alcoholic cirrhosis [\[42](#page-11-31)]. This fact is worrying because it tells us a lot about the role that addictions can play after bariatric surgery [\[43\]](#page-11-32); furthermore, it may play a determining role in long-term weight loss, which should be also taken into account in future studies on these patients.

It's imporant to emphasize that this analysis involved screening 1126 studies, with 1032 records reviewed and 247 assessed for eligibility. Despite identifying 16 studies with the necessary data, 140 studies lacked data stratifed by liver status at baseline, revealing a common issue of incomplete data reporting. Efforts to contact original authors resulted in a low response rate (12%), with only 13 out of 108 authors responding and just seven providing the needed data. These challenges highlight the need for standardized reporting practices and better data sharing to enhance the quality of systematic reviews and meta-analyses [[22](#page-11-14)].

## **Limitations**

Several studies report signifcant diferences in age or BMI between groups. This variability could affect the generalizability of fndings and emphasizes the need to control for these factors in analyses. Additionally, diferences in sex distribution  $(>15\%$  difference) in some studies might influence weight loss outcomes, as gender can afect metabolic results and weight loss patterns. Most studies used liver biopsy for diagnosis that supports comparability, but diferent classifcations used (NAS, METAVIR, and SAF Score) might infuence the results. The limited number of studies could also contribute to variability and heterogeneity in the results.

# **Conclusions**

While MASLD does not appear to significantly affect shortterm weight loss outcomes, there is notable variability in long-term results, emphasizing the need for standardized reporting practices in future research. Specifcally, in the context of SG, patients with MASH tend to experience a reduced degree of weight loss compared to those without MASH. Future studies should focus on complete and transparent data reporting to facilitate meta-analyses and systematic reviews. Establishing a culture of cooperation and transparency within the research community is crucial for enhancing the quality and credibility of results, leading to more accurate and generalizable conclusions.

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**Data Availability** The data supporting the fndings of this study are available within the publication and upon reasonable request to the corresponding author.

## **Declarations**

**Competing Interest** The authors declare no competing interests.

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