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A Multidisciplinary Approach for Nonresponders Following Bariatric Surgery: What Is the Value?

Marleen M. Romeijn, MD,^{1,2,*} Martine Uittenbogaart, MD,¹ François M.H. van Dielen, MD, PhD,¹
Arijan A.P.M. Luijten, MD,¹ Loes Janssen, PhD,¹ and Wouter K.G. Leclercq, MD¹

Background: Approximately 25% of patients after bariatric surgery either do not lose enough weight or regain a considerable amount of weight, both are referred to as nonresponse. This study aimed to describe the added value of a multidisciplinary approach on treatment strategies in patients with nonresponse.

Materials and Methods: The primary outcome of this retrospective cohort study was the initiated treatment by the multidisciplinary team (MDT). Outcomes were described separately for patients with primary (i.e., <50% excess weight loss [EWL]) and secondary nonresponse (i.e., ≥50% EWL followed by >5% regain).

Results: Of the 83 included patients, 10 patients underwent revisional surgery. A total of 73 patients received a conservative treatment as they either had not been able to change their lifestyle or due to certain behavioral factors. Conservatively treated patients stabilized in weight after 2 years ($-0.9 \text{ kg} \pm 5.8$, $n = 27$), while surgically treated patients did lose weight ($-12.1 \text{ kg} \pm 16.9$, $n = 7$). One patient suffered from an ulcerative stenosis at the gastroenterostomy after limb length alteration.

Conclusions: a conservative treatment was the most frequently advocated treatment by the MDT. A surgical treatment resulted in successful weight loss, although only a few patients were selected for this by the MDT. A multidisciplinary approach can be beneficial for the identification of lifestyle and behavioral factors.

Keywords: bariatric surgery, obesity, weight regain, multidisciplinary team, revisional surgery

Introduction

BARIATRIC SURGERY HAS proven to be the most effective treatment for obesity to achieve long-term substantial weight loss.^{1,2} However, ~25% of patients do not respond well as they lose an insufficient amount of weight, or regain weight after sufficient weight loss.^{3,4} These phenomena can be described as primary nonresponse and secondary nonresponse. One can distinguish between these two types of nonresponse by applying a cutoff score. Frequently used cutoff scores are the amount of excess weight loss (EWL) and the amount of total weight loss (TWL) reported 1 to 2 years postsurgery.^{5–7} The etiology for nonresponse can be lifestyle related, but can also be explained by anatomical alterations like a dilated pouch, dilated sleeve, dilated gastroenterostomy, or fistula formation such as gastrogastroic fistula.^{8,9}

Revisional bariatric surgery constitutes a popular solution for patients with nonresponse.¹⁰ Commonly offered procedures are conversion from sleeve gastrectomy (SG) to Roux-en-Y gastric bypass (RYGB), one anastomosis gastric bypass/mini gastric bypass (OAGB/MGB), lengthening of biliopancreatic limb, band, or ring placement, revision of the gastric pouch and/or stoma, and a biliopancreatic diversion/duodenal switch (BPD/DS).^{7–13} To date, the long-term efficacy and safety of these procedures is difficult to predict. Lengthening of the biliopancreatic limb may result in severe malnutrition leading to the need for reoperation, whereas ring placement may result in reoperation due to dysphagia and/or ring migration.^{8,13}

In 1991, the National Institutes of Health Consensus Statement advocated that a multidisciplinary team (MDT) is required to optimize bariatric patient care.¹⁴ This resulted in the formation of dedicated teams consisting of bariatric surgeons,

¹Department of Surgery, Máxima Medical Center, Veldhoven, The Netherlands.

²Research School NUTRIM, Department of Surgery, Maastricht University Medical Center, Maastricht, The Netherlands.

*ORCID ID (<https://orcid.org/0000-0001-6582-5255>).

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obesity physicians, dieticians, physical therapists, and medical psychologists. The beneficial effect of MDTs on surgical outcomes has extensively been described in the field of surgical oncology^{15–17}; however, the effect of a MDT on nonresponse after bariatric surgery is relatively poorly understood. It was Srivastava and Buffington who demonstrated that an intensive multidisciplinary lifestyle intervention combined with medication could improve early weight loss in patients with nonresponse.¹⁸ This suggests that the role of a MDT needs further expansion in the management of nonresponders after bariatric surgery and triggers the need for a more extensive analysis. The aim of this study was to describe the added value of a multidisciplinary approach, in a bariatric tertiary referral center, on treatment strategy in nonresponders after bariatric surgery.

Materials and Methods

Study population

Data about nonresponders after RYGB and SG was collected in our center from the first of January 2016 till the first of December 2020. Patients were included if they regained $\geq 5\%$ weight with respect to the lowest postoperative weight after RYGB or SG (i.e., nadir weight). At the moment of inclusion, $\geq 5\%$ regain was a commonly used cutoff score.⁶ Patients were excluded in case of a banded RYGB and OAGB/MGB because these were less frequently performed procedures and this would have introduced heterogeneity otherwise. A history of laparoscopic adjustable gastric banding or Mason gastroplasty before the RYGB or SG was no criteria for exclusion. Ethical approval was obtained from the Medical Ethics Committee of our center, reference number: N19.054 (L19.065). For this type of study, formal consent from all individual participants was not required.

The primary procedure could have taken place in our center or elsewhere. In our center, all patients were screened for primary bariatric surgery by a MDT, and IFSO criteria were used for qualification.¹⁹ Patients were monitored in the outpatient clinic by members of the MDT for a period of 5 years. Patients were categorized as primary nonresponders if the EWL was $< 50\%$ after primary surgery; patients were categorized as secondary nonresponders if the EWL exceeded the 50% EWL threshold and a regain of $> 5\%$ was reported as per nadir weight. The below section elaborates on the process and approach taken before and during the multidisciplinary assessment.

Assessment before MDT meeting

Initial assessment of the patient was done by a bariatric surgeon. Subsequently, the patient was referred to a dietician and physical therapist for assessment of nutritional habits and physical activity. The dietician focused on food intake, food choices, feelings of satiety, and hunger and signs of emotional eating. The physical therapist focused on activity habits. If indicated, consultation by a medical psychologist was offered.

MDT meeting

Following individual consultation by all team members, the treatment strategy was discussed in a joint multidisciplinary meeting with weekly occurrence. At least one member of

the following fields of expertise attended this meeting: bariatric surgeon (chair), nurse practitioner, dietician, physical therapist, and medical psychologist. Notably, under certain circumstances, the patient was discussed in the meeting despite that this patient was not seen by all team members (e.g., persistent lack of attendance at one of the appointments).

Treatment strategy

After evaluation by the MDT, a decision was taken whether the patient needed a lifestyle and/or behavioral intervention or whether the patient was qualified for revisional surgery. For the purpose of this study, treatment options were divided into conservative and surgical treatment (Fig. 1). A conservative treatment consisted of a nutritional and/or physical intervention, summarized as “lifestyle.” A nutritional intervention was indicated, for example, in case of unhealthy food choices and detrimental eating patterns, whereas a physical intervention was indicated in case of a sedentary lifestyle with the goal to increase activity habits. Moreover, a behavioral intervention was indicated if there were signs of emotional eating and problems in impulse control. If there were signs indicating an eating disorder, patients were referred to a psychiatric clinic specialized in treatment of such disorders.

The indication for revisional surgery was not based on the degree of weight loss or regain, so no cutoff scores were applied by any surgeon. The type of procedure depended on the index procedure, perioperative findings, and expert opinion. Preoperative upper gastrointestinal series and upper endoscopy were performed in patients who were offered revisional surgery.

Study outcomes

The primary outcome was the initiated treatment by the MDT, either conservatively or surgically. Secondary outcomes were weight loss achieved in a period of 4 years after start of the treatment and complications after surgical treatment. Weight loss was described as %EWL, which can be calculated as follows: $(\text{preoperative weight} - \text{nadir weight}) / (\text{preoperative weight} - \text{ideal body weight}) \times 100\%$. Ideal body weight was based on a body mass index (BMI) of 25 kg/m^2 ; initial body weight was the weight at the moment of screening. Weight loss was also expressed in %TWL, which can be calculated as follows: $(\text{preoperative weight} - \text{nadir weight}) / \text{preoperative weight} \times 100\%$, or $(\text{postoperative highest weight} - \text{postoperative lowest weight achieved after treatment of nonresponse}) / \text{postoperative highest weight} \times 100\%$. The percentage of regain was calculated as percentage kg gained after reaching the lowest postoperative weight (nadir).

Statistical analyses

Quantitative data are presented as mean with standard deviation (range) or median with interquartile range; categorical data are expressed in numbers and percentages. A paired *t*-test was performed to compare the weight before and after the treatment of nonresponse initiated by the MDT. All analyses were performed using the program Statistical Package for Social Sciences version number 22.0 (IBM SPSS 22.0; Chicago, IL).

FIG. 1. Flowchart of multidisciplinary management in patients with nonresponse after bariatric surgery. MDT, multidisciplinary team.

Results

Description of the population

A total of 119 patients were initially included in the study. Fifteen patients were excluded due to either having a banded RYGB ($n=9$), an OAGB/MGB ($n=2$), or because they did not receive RYGB or SG but a different procedure (i.e., laparoscopic adjustable gastric banding $n=3$, BPD $n=1$). Of the 104 remaining patients, 21 patients did not attend their appointment with the dietician, physical therapist, and/or medical psychologist. Therefore, no optimal treatment could be advised and, consequently, these patients were excluded from further analysis. All 83 remaining patients were referred, before the multidisciplinary meeting, to a dietician and a physical therapist. Twenty-six patients (31.3%) were referred to a medical psychologist as well.

The baseline characteristics of the 83 included patients are demonstrated in Table 1. The group of conservatively treated patients consisted of 73 patients (88%), whereas the group of surgically treated patients consisted of 10 patients (12%). There were no significant differences in terms of gender, preoperative BMI, and percentage of regain between the two groups. Based on the aforementioned definitions, 14 patients (16.9%) were defined as primary nonresponders, who received a conservative treatment. The remaining 69 patients (83.1%) were defined as secondary nonresponders and within this group, 59 patients received a conservative treatment, and 10 patients a surgical treatment.

When focusing on the conservative treatment, 63 patients (86%) received a nutritional intervention, 46 patients (63%) a physical intervention, and 18 patients (25%) a behavioral intervention. When focusing on the surgical treatment, four different procedures were performed as described in Table 2. The index procedure was RYGB in eight patients and SG in two patients. In particular, two patients with prior SG underwent conversion to RYGB; three patients with perioperative common limb lengths between 440 and 600 cm underwent

shortening of the common limb to 200–380 cm; four patients with a perioperative large pouch and/or stoma underwent re-sizing; and one patient received a gastric ring (Minimizer).

Weight loss outcomes

Table 3 displays the effect of the treatment, initiated by the MDT up to 4 years after start of the treatment. As result of a conservative treatment, patients lost on average 0.9 kg within 2 years ($SD=5.8$) and gained 0.2 kg within 4 years ($SD=9.4$). Recalculated in %TWL, the weight loss was 1.2% ($SD=5.5$), 0.7% ($SD=6.3$), 2.7% ($SD=10.5$), and 0% ($SD=9.3$) within, respectively 1, 2, 3, and 4 years. Additional analysis showed that 4 years after start of the conservative treatment ($n=17$) 47% lost weight, while 53% gained weight. As result of a surgical treatment, patients lost on average 12.1 kg within 2 years ($SD=16.9$) and even more within 3 and 4 years; however, only a small number of patients was assessed at these time points. The weight loss 1, 2, and 3 years after revisional surgery was statistically significant ($p \leq 0.01$). Recalculated in %TWL, the weight loss was 12.1% ($SD=12.0$), 11.6% ($SD=15.8$), 26.9% ($SD=0.6$), and 23.9% ($n=1$) within, respectively, 1, 2, 3, and 4 years.

Complications after surgical treatment

One postoperative complication occurred in a patient with a history of gastric banding, followed by RYGB and subsequent alteration of the limb length. This patient developed an ulcerative stenosis at the gastroenterostomy 1 year after alteration, presumably due to nonsteroidal anti-inflammatory drugs use and the presence of *Helicobacter pylori*. The patient was successfully treated with endoscopic dilatation and received eradication therapy for this *H. pylori*.

Lost to follow-up

After 2 years, 49 patients (59.1%) were lost to follow-up, and after 4 years, this number increased to 65 patients

TABLE 1. BASELINE CHARACTERISTICS OF THE STUDY POPULATION

	Total, n = 83	Conservative treatment, n = 73	Surgical treatment, n = 10
Gender, n (%)			
Female	75 (90.4)	66 (90.4)	9 (90.0)
Age at the time of MDT meeting (years)	43.7 ± 9.8 (41.6–45.8)	44.0 ± 9.9 (41.7–46.3)	41.4 ± 9.2 (34.9–47.9)
Preoperative comorbidities, n (%)			
Hypertension	17 (20.5)	14 (19.2)	3 (30.0)
Type II diabetes	7 (8.4)	5 (6.8)	2 (20.0)
Dyslipidemia	4 (4.8)	4 (5.5)	—
OSAS	7 (8.4)	7 (9.6)	—
Osteoarthritis	5 (6.0)	5 (6.8)	—
History of bariatric surgery, n (%)			
LAGB	17 (17.5)	14 (19.2)	3 (30.3)
Mason	6 (7.2)	6 (8.5)	—
SG	5 (6.0)	5 (6.8)	—
Index procedure, n (%)			
RYGB	73 (88.0)	65 (89.0)	8 (80.0)
SG	10 (12.0)	8 (11.0)	2 (20.0)
Laparoscopy	79 (95.2)	69 (94.5)	10 (100.0)
Complications, n (%)			
No complications	73 (88.0)	63 (86.3)	10 (100.0)
Clavien-Dindo grade I	1 (1.2)	1 (1.4)	—
Clavien-Dindo grade II	2 (2.4)	2 (2.7)	—
Clavien-Dindo grade III	7 (8.4)	7 (9.6)	—
Preoperative weight (kg)	127.6 ± 20.7 (123.0–132.1)	126.2 ± 20.3 (123.0–132.1)	137.5 ± 21.6 (122.1–152.9)
Preoperative BMI (kg/m ²)	44.7 ± 7.0 (43.2–46.2)	44.7 ± 7.4 (43.1–46.2)	44.9 ± 3.5 (42.4–47.5)
Maximal EWL (%)	77.9 ± 23.7 (72.8–83.2)	78.6 ± 25.0 (72.8–83.2)	73.7 ± 10.1 (66.4–80.9)
Maximal TWL (%)	32.9 ± 9.7 (30.8–35.0)	32.9 ± 10.2 (30.8–35.0)	32.2 ± 3.6 (29.7–34.8)
Amount of primary nonresponse (%)	14 (16.9)	14 (19.2)	—
Amount of secondary nonresponse (%)	69 (83.1)	59 (80.8)	10 (100.0)
Weight regain (%)	24.9 ± 15.1 (21.6–28.2)	25.5 ± 15.7 (21.6–28.2)	20.9 ± 9.1 (14.4–27.5)
Interval to presentation (years)	4.1 ± 2.2 (3.6–4.6)	4.0 ± 2.2 (3.6–4.6)	5.0 ± 1.8 (3.7–6.3)
Analysis of nonresponse			
Upper gastrointestinal series (%)	28 (33.7)	18 (24.7)	10 (100.0)
Endoscopy (%)	5 (6.0)	3 (4.1)	2 (20.0)

Data presented as n (%), mean with standard deviation and 95% confidence interval. Primary nonresponse is defined as <50% EWL. Secondary nonresponse is defined as ≥50% and a regain of ≥5% with respect to nadir weight.

BMI, body mass index; EWL, excess weight loss; LAGB, laparoscopic adjustable gastric banding; MDT, multidisciplinary team; OSAS, obstructive sleep apnoea syndrome; RYGB, Roux-en-Y gastric bypass; SG, sleeve gastrectomy; TWL, total weight loss.

TABLE 2. OVERVIEW OF GIVEN TREATMENT FOR NONRESPONSE, INITIATED BY THE MULTIDISCIPLINARY TEAM

Total, n = 83	
Conservative treatment n = 73, n (%)	
Nutritional intervention	63 (86.3)
Physical intervention	46 (63.0)
Behavioral intervention	18 (24.7)
Referred to psychiatric institution	4 (5.5)
Surgical treatment n = 10, n (%)	
Alternation of the limb length	3 (30.0)
Revision of the gastric pouch and/or stoma	4 (40.0)
RYGB	2 (20.0)
Ring placement	1 (10.0)

Data presented as n (%).

(78.3%). The reason of lost to follow-up is unknown. The baseline characteristics were analyzed comparing missing patients with nonmissing patients 2 years after the start of the treatment. This analysis showed that the percentage of regain was significantly higher in the group of missing patients (28.3% vs. 20.3%, $p = 0.02$). On the contrary, the prevalence of preoperative hypertension and type 2 diabetes was lower in the group of missing patients (10.4% vs. 34.3%, $p = 0.01$ and 2.1% vs. 17.1%, $p = 0.02$, respectively). Other baseline characteristics did not differ between the two groups.

Discussion

There is no standardized approach in the treatment of nonresponders following bariatric surgery. While ongoing studies about the efficiency of revisional surgery are being reported in literature,^{8,9,11–13} studies about the effect of a

TABLE 3. EFFECT OF A CONSERVATIVE AND SURGICAL TREATMENT ON WEIGHT IN NONRESPONDERS FOLLOWING BARIATRIC SURGERY

	<i>Conservative treatment</i>				<i>Surgical treatment</i>			
	<i>Primary nonresponse</i>		<i>Secondary nonresponse</i>		<i>Primary nonresponse</i>		<i>Secondary nonresponse</i>	
	<i>Total</i>				<i>Total</i>			
Weight at start of treatment (kg)	104.8±20.9 (99.9 to 109.7)	n=73	123.0±24.3 (109.0 to 137.1)	n=14	100.5±17.6 (95.9 to 105.0)	n=59		
ΔWeight (kg's) ≤1 year	-1.7±6.1 (-3.7 to +0.3)	n=39	p=0.09	-0.9±2.9 (-3.3 to -1.5)	n=8	p=0.39	-1.9±6.8 (-4.3 to +0.6)	n=31
ΔWeight (kg's) ≤2 years	-0.9±5.8 (-3.2 to +1.4)	n=27	p=0.44	-3.9±4.7 (-11.5 to +3.4)	n=4	p=0.19	-0.3±5.9 (-2.9 to +2.2)	n=23
ΔWeight (kg's) ≤3 years	-3.4±10.8 (-8.0 to +1.2)	n=24	p=0.14	-5.2±11.9 (-19.9 to +9.5)	n=5	p=0.38	-2.9±10.9 (-8.2 to +2.3)	n=19
ΔWeight (kg's) ≤4 years	+0.2±9.4 (-4.7 to +5.0)	n=17	p=0.92	-0.04±7.6 (-9.5 to +9.4)	n=5	p=0.99	+0.4±10.3 (-6.2 to +6.9)	n=12
								p=0.91
<i>Surgical treatment</i>								
Weight at start of treatment (kg)	115.6±17.9 (102.8 to 128.4)	n=10		×	115.6±17.9 (102.8 to 128.4)	n=10		
ΔWeight (kg's) ≤1 year	-13.0±12.1 (-22.3 to -3.7)	n=9	p=0.001 ^a	×	-13.0±12.1 (-22.3 to -3.7)	n=9	p=0.001 ^a	
ΔWeight (kg's) ≤2 years	-12.1±16.9 (-27.7 to +3.5)	n=7	p=0.003 ^a	×	-12.1±16.9 (-27.7 to +3.5)	n=7	p=0.003 ^a	
ΔWeight (kg's) ≤3 years	-26.3±0.4 (-29.4 to -23.1)	n=2	p=0.000 ^a	×	-26.3±0.4 (-29.4 to -23.1)	n=2	p=0.000 ^a	
ΔWeight (kg's) ≤4 years	-27.0	n=1	×	×	-27.0	n=1	×	

Data presented as mean with standard deviation and 95% confidence interval.

^aPaired *t*-test: significant difference compared to the weight at the start of the treatment, $p \leq 0.05$.

multidisciplinary approach fall behind. Although it is proposed that a multidisciplinary approach is useful in the analysis of patients with nonresponse,^{7,9,13,20} this study is the first in describing the added value of such a multidisciplinary approach.

The main finding of this study was that the MDT advocated a conservative treatment more frequently than a surgical treatment (88% vs. 12%). The majority of patients were considered not to be eligible for revisional surgery as they either had not been able to change their lifestyle or due to certain behavioral factors. Based on previous research in our center between 2012 and 2015, it has been identified that before the introduction of a MDT, 68% of the nonresponders underwent surgery after failed RYGB.¹¹ That study included 65 patients with weight loss failure and weight regain who were consulted by a bariatric surgeon. Consultation by a dietitian and physical therapist and/or medical psychologist was not scheduled on a routine basis. Furthermore, no joint meeting by the MDT took place. The result of this study (68% revisional surgery) is in sharp contrast with the result of the current study (12% revisional surgery). One can argue that the MDT may have underselected appropriate candidates for revisional surgery or alternatively may have optimized the selection of candidates. The small sample size and heterogeneity in revisional procedures in the current study restricts making firm statements about this.

When reviewing weight loss outcomes, the weight loss achieved by revisional surgery was superior to the weight loss achieved by a conservative treatment. Notably, one could also argue that the weight was sustained as result of a conservative treatment, indicating that it may prevent patients from gaining more weight in the future. The question arises why a conservative treatment should be advocated as this resulted in less weight loss. In the context of revisional surgery it is worth noting that a patient is exposed to the risks of major complications and the risk of unsustainable weight loss in the long term.^{8,21} A reason for this can be that surgery does not resolve the lifestyle and/or behavioral problems that may have contributed to nonresponse in the first place.^{20,22,23}

The observed complication rate within our study is in line with other literature. Complication rate after pouch/anastomosis revision is estimated at 3.5% and after limb alteration at 12%.⁸ Revision of the gastric pouch and/or stoma was most frequently performed in this study. The four patients who underwent this procedure did not suffer from any complications. From the three patients who underwent limb alternation, one patient suffered from an ulcerative stenosis at the gastroenterostomy, which is a frequently reported complication in literature.^{24,25}

This study distinguished outcomes between primary and secondary nonresponders. Previous studies did not report treatment outcomes using this classification, although it is proposed as standardized terminology.^{6,10} This study supports this classification as it was found that patients with secondary nonresponse rather than primary nonresponse were treated surgically. This may indicate the need for a different treatment strategy in patients with primary and secondary nonresponse. A note of caution might be appropriate as there were demographical differences between the two groups. Speculatively, primary nonresponse might be a manifestation of insufficiently treated or even untreated eating behaviors and/or psychological problems, whereas

TABLE 4. CHECKLIST OF FACTORS THAT CAN BE USED DURING THE ASSESSMENT OF A PATIENT WITH NONRESPONSE ADVOCATING A CONSERVATIVE OR SURGICAL TREATMENT

<i>Factors in favor of conservative treatment</i>	<i>Factors in favor of surgical treatment</i>
Excessive dietary intake	Signs of surgical failure, e.g., pouch or stoma dilatation,
Inappropriate food choices	gastrogastric fistula
Lack of satiety	Dietary compliance
Constant hunger sensation	No signs of maladaptive eating habits
Emotional eating	Physically active
Binge eating	No signs of mental health disorder
Sweet eating	No signs of alcohol or drug use
Grazing behavior	
Physically inactive	
Mental health disorder, e.g., depression, anxiety, personality disorders	
Alcohol or drug use	

secondary nonresponse might indicate anatomical problems. This theory warrants further research to obtain more information about the etiology and treatment strategy that is most efficient per nonresponse type.

This retrospective study has several limitations that should be considered. Not all patients received their primary surgery in our center and therefore information about preoperative screening and treatment could have been missed. It is questionable whether certain lifestyle and/or behavioral factors were present before primary surgery and whether the patients' adjustments met the standards used in our center. Another limitation could be that the MDT experienced a learning curve as the multidisciplinary evaluation of nonresponders was introduced in 2016 in our center. It is possible that the different team members experienced a change in their evaluation and subsequent treatment. Moreover, pharmacological therapy was not used, while this has shown beneficial results in the treatment of nonresponse.²⁶ It should also be mentioned that results regarding weight loss and complication rates are limited due to the small selected group of patients who underwent revisional surgery. Furthermore, the 20% of patients who were excluded from the analysis due to no-show may have introduced selection bias, as well as the patients who did not attend their follow-up appointments after the treatment was set. The latter may be explained by further weight regain or by a loss of confidence in the MDT.²⁷ Notwithstanding these limitations, the study offers insights into the treatment of nonresponders after bariatric surgery. For further practices, a checklist of relevant factors was made proposing a conservative or surgical treatment in nonresponders (Table 4).

Conclusion

This study demonstrated that when patients with nonresponse after bariatric surgery are evaluated by a MDT, a conservative treatment is more often applied than a surgical treatment. The MDT identified patient-specific factors presumably contributing to the nonresponse and these factors constituted the framework for a conservative treatment. At present, the question that remains unanswered is whether a

multidisciplinary approach set the right treatment as perhaps the selection of candidates for revisional surgery was incorrect. More studies are required to position the multidisciplinary approach in the treatment of nonresponse, as well as determine what kind of revisional procedure should be preferred.

Authors' Contributions

M.R., L.J., F.D., and W.L., conceived the original study and design. M.R. is responsible for data collection, data recording, and preparing the article. M.U. and A.L., contributed to writing the article. All authors have read and approved the final article.

Author Disclosure Statement

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Address correspondence to:

Marleen M. Romeijn, MD
Research School NUTRIM
Department of Surgery
Maastricht University Medical Center
Universiteitssingel 40
Maastricht 6229 ER
The Netherlands

E-mail: m.romeijn@maastrichtuniversity.nl;
bariatric.resurge@mmc.nl