

# Approach to Weight Loss Patient's Post Bariatric Surgery

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**Abstract:** *Approach to obesity management includes lifestyle changes, pharmacotherapy, and bariatric surgery when indicated. Bariatric surgery is one of the most effective tools for achieving clinically significant weight reduction, with different procedures leading to varying degree of weight loss. Bariatric surgery is a safe and effective option for those who do not achieve durable weight loss with non - surgical methods. There is a potential role of pharmacotherapy in combination with bariatric surgery to prevent weight recurrence.*

**Keywords:** obesity, weight, bariatric surgery, body mass index (BMI)

## 1. Introduction

Obesity is defined by the World Health Organization (WHO) as an abnormal or excessive fat accumulation that presents a risk to health, commonly classified by the body mass index (BMI): Underweight < 18.5%; normal weight 18.5% - 24.9%; overweight 25% - 29.9%; obese class I 30% - 34.9%; obese class II 35% - 39.9%; obese class III >40 %.<sup>(1 - 4)</sup> BMI is simple to calculate, but it does have its limitations where factors such as age, muscle mass and ethnicity can influence its relationship with body fat.<sup>(1 - 4)</sup> Other measures such as skinfold thickness, waist circumference and waist - to - hip ratio are increasingly used to assess an individual's risk of obesity related conditions such as type 2 diabetes mellitus and cardiovascular disease.<sup>(1 - 4)</sup> In response to the expanding weight problem, there has been an increased focus on weight loss for improving clinical outcomes in morbidly obese patients. Major medical disorders that are directly induced or exacerbated by obesity.<sup>(5, 6)</sup> Obesity is associated with osteoarthritis, insulin - resistant diabetes mellitus, obstructive sleep apnea, dyslipidemia and secondary coronary artery disease, hypertension and a group of miscellaneous disorders, including gastroesophageal reflux disease, cardiovascular disease, asthma and deep vein thrombosis, increased risk of endometrial, breast, pancreatic, prostate and colon cancer.<sup>(5, 6)</sup> Unfortunately, the average weight loss reported to be achieved by medical weight loss programs is neither sufficient nor durable enough to be an adequate treatment for morbid obesity.<sup>(7, 8)</sup> According to WHO, obesity has reached epidemic proportions worldwide, with approximately 1.9 billion overweight and 650 million obese adults.<sup>(9 - 11)</sup> Purnell explains the physiology of body weight regulation, focusing on the communication between the brain, gastrointestinal system, and fat tissue. The dysregulation of this system can lead to obesity and its associated with health complication.<sup>(9)</sup>

Obesity is a complex, multifactorial disease that occurs through interaction between genetic predisposition and the environment. The genetic contributes to weight through syndromic, monogenic, and polygenic causes. The leptin - melanocortin hypothalamic pathway is a core mechanistic pathway regulating appetite. Environmental changes (ultra -

processed food, sleep changes, medications, parental exposures) has driven the more recent obesity epidemic.<sup>(9 - 12)</sup> Bariatric surgical procedures reduce caloric intake by modifying the anatomy of the gastrointestinal tract. These operations are classified as either restrictive or malabsorptive. Restrictive procedures limit intake by creating a small gastric reservoir with a narrow outlet to delay emptying. Malabsorptive procedures bypass vary.<sup>(13)</sup>

## 2. Updates on surgical and procedural interventions for obesity

### 1) Sleeve gastrectomy (SG)

This restrictive procedure involves resection of a large segment of the stomach, leaving a narrow gastric remnant and intact segments of the intestine. Resection of the gastric fundus restricts the ability of the residual stomach to stretch and nearly eliminates the production of ghrelin (a foregut peptide hormone implicated in regulating food intake and body weight), and a reduce a risk of death.<sup>(14 - 19)</sup> SG can yield excess weight loss of up to 70% within one year, which is maintained to at least three years.<sup>(20)</sup> SG increases remission rates of diabetes (86%), hypertension (82%), dyslipidemia (83%) and sleep apnea (91%). SG has a low mortality rate and its most serious complication is leakage, which occurs in 2-3% of patients.<sup>(21)</sup> Others include strictures (4%) and gastro - oesophageal reflux.<sup>(22)</sup>

### 2) Roux - en - Y gastric bypass (RYGB)

This procedure combines restrictive and malabsorptive components to facilitate weight loss. The stomach is restricted to a small pouch and the small intestine is divided and rerouted; the jejunum is anastomosed to the small gastric pouch (the Roux limb), whereas the proximal small intestine (including the duodenum) is further anastomosed distally, in the jejunum (the biliopancreatic limb). Food from the gastric pouch travels through the Roux limb into the common channel, where it mixes with biliopancreatic secretions for digestion; the shorter the common channel, the greater the malabsorptive effect.<sup>(14 - 19)</sup>

(RYGB) estimated weight loss of 73% within 1 - year, good long - term weight loss maintenance, and remission of comorbidities such as diabetes (95%), dyslipidaemia (80%), hypertension (81%) and sleep apnoea (95%).<sup>(20)</sup> 30 - day mortality rates are lower for the laparoscopic vs open approach (0.2% vs 2.1%, respectively).<sup>(23)</sup> Serious complications include anastomotic leaks (3–5%) and internal bowel herniation (3.1%) with the potential to lead to bowel obstruction and perforation.<sup>(24 - 26)</sup> Other complication includes malabsorption of dietary vitamins (particularly B - complex vitamin) and minerals. The vitamin B12 deficiency may induce myelopathy, degenerative of the posterior and lateral column and macrocytic anemia.<sup>(14 - 19)</sup> Thiamine deficiency can be induced by malabsorptive bariatric surgical procedures. It can also be seen in conjunction with excessive alcohol use and exists worldwide as consequence of major nutritional deficiency. The classic presentation of thiamine deficiency includes ataxia, nystagmus, and confusion (Wernicke's encephalopathy). Patient with thiamine deficiency may have distal sensorimotor polyneuropathy (Dry Beriberi) with or without cardiomyopathy (Wet Beriberi) and amnesia with confabulation but relatively normal cognition (Korsakoff's syndrome).<sup>(14 - 19)</sup> Prompt identification of thiamine deficiency in cases of Wernicke's encephalopathy and beriberi is critical; the rapid administration of intravenous thiamine may restore normal neurologic function and prevent irreversible damage. Investigation for other nutritional deficiencies should also be conducted.<sup>(14 - 19)</sup>

### 3) Adjustable Gastric Banding (ABS)

This purely restrictive procedure involves the placement of an adjustable band around the gastric entry to partition the stomach into two compartments. The adjustable band can be tightened or relaxed through an injectable saline - infusion port that is placed in the subcutaneous tissue.<sup>(14 - 19)</sup> Longitudinal assessment of bariatric surgery yield excess weight loss of up to 55% 2 years post - operatively, but also promotes the remission of diabetes (74%), hypertension (54%), dyslipidaemia (40%) and sleep apnoea (94%).<sup>(23)</sup> The Longitudinal Assessment of Bariatric Surgery consortium have reported no mortality at 30 days post - LABG (n=1, 198).<sup>(23)</sup> More common minor complications include oesophageal pouch dilatation (11%) and gastroesophageal reflux, whereas major complications include band slippage (7.9%) and erosion (<1%).<sup>(27 - 29)</sup>

### 4) Biliopancreatic diversion with a duodenal switch (BPD - DS)

This procedure includes a restrictive component (a sleeve gastrectomy), but it is primarily a malabsorptive procedure reserved for patients with severe obesity. A gastroileostomy with a very short common channel is constructed to maximize malabsorption; the mixture of food contents and biliopancreatic secretions is diverted to the distal segments of the small intestine.<sup>(14 - 19)</sup> BPD - DS Can yield excess weight loss of up to up to 73% with maintenance to over 8 years via both restrictive and malabsorptive mechanisms akin to RYGB, with significant effects on comorbidities such as type2 diabetes.<sup>(28)</sup> Although mortality rates remain low (<1%), BPD - DS is a more complex procedure, with complications including perioperative anastomotic leaks (3–4%) and splenectomy (<1%), and later malnutrition (4%),

internal bowel herniation and small bowel obstruction (2 - 7%).<sup>(30, 31)</sup>

### Post - metabolic bariatric surgery and type 2 diabetes mellitus remission

Metabolic surgery is highly effective in obtaining significant improved quality of life and a reduced risk for other obesity - related and diabetes - related disorders, including microvascular disease, sleep apnea, fatty liver disease, and malignant diseases.<sup>(32 - 35)</sup> Comparable glycemic benefits of surgery have been observed in patients with lower initial BMIs <35 kg/m<sup>2</sup>. In the STAMPEDE trial, the antidiabetic effects of surgery were the same in patients with baseline BMIs higher and lower than 35 kg/m<sup>2</sup>.<sup>(36 - 38)</sup>

Shah et al.<sup>(39)</sup> demonstrated a substantial benefit of RYGB in Indian patients with an initial BMI <35 kg/m<sup>2</sup>, and similar BMI - independent benefits of surgery have been established in meta - analyses.<sup>(39 - 41)</sup> As a result, current guidelines support the use of metabolic surgery in individuals with mild obesity and poorly controlled type 2 diabetes.<sup>(41 - 44)</sup> Although the antidiabetic effects of ABS appear less powerful than those of other metabolic operations, one study nonetheless demonstrated the ability of ABS to induce remission of type 2 diabetes mellitus, defined as fasting glucose <126 mg/dl and post - load glucose during a 2 - h oral glucose tolerance test <200 mg/dl, at least 2 days after cessation of antidiabetic medication, among overweight (BMI 25 to 30 kg/m<sup>2</sup>) individuals.<sup>(45)</sup> In some patients, the bariatric and metabolic effects of gastrointestinal surgery may diminish over time, with relapse of type 2 diabetes in patients who had initially achieved full remission. Even in these patients, however, the recurrent diabetes is generally far less severe, requires less intensive medication regimens, and is more easily controlled than before the metabolic surgery. Patients who are older, use insulin, regain weight, or have greater baseline waist circumference, longer diabetes duration, or poor preoperative glycemic control appear to be at highest risk for late postoperative diabetes recurrence.<sup>(46 - 48)</sup> Even in the setting of diabetes relapse, however, a legacy effect may exist. Finally, the effect of post - operative antidiabetic medications in patients whose diabetes failed to remit or recurred late after surgery is currently being examined (Canagliflozin vs. Placebo for Post Bariatric Patients with Persistent Type 2 Diabetes [CARAT]).<sup>(49)</sup> A courcoulas et al, demonstrate prospective randomized interventional data from the largest cohort of patients to date demonstrates that metabolic surgery improves glycemic control, diabetes - related comorbidities, and weight loss to a greater extent than medical/lifestyle intervention for up to three years after treatment, with minimal and generally tolerable adverse events.<sup>(50)</sup> The study show that surgery is more effective than medical therapy in the long - term control of type 2 diabetes. Our findings suggest that it might be possible to reduce the frequency of glycemic monitoring in patients who maintain at least five years of remission of type 2 diabetes although larger studies are needed to confirm this finding. Clinicians and policy makers should ensure that metabolic surgery is appropriately considered in the management of patients with obesity and type 2 diabetes.<sup>(51)</sup> A very long - term follow - up observational study of obese patients with type 2 diabetes, bariatric surgery was associated with more frequent diabetes remission and fewer complications than usual care. These

findings require confirmation in randomized trials.<sup>(52)</sup> Weight loss after bariatric surgery is strongly associated with initial Type 2 Diabetes remission; however, above a threshold of 20% total weight loss, rates of initial Type 2 Diabetes remission did not increase substantially. Achieving this threshold is also associated with initial remission even in patients who traditionally experience lower rates of remission, such as patients taking insulin.<sup>(53)</sup>

### Post - metabolic bariatric surgery weight loss and prevalence of nonalcoholic steatohepatitis (NASH) remission

Bariatric - metabolic surgery was more effective than lifestyle intervention and best medical care as a treatment of nonalcoholic steatohepatitis in people with obesity, with or without type 2 diabetes. Roux - en - Y gastric bypass and sleeve gastrectomy had similar efficacy on nonalcoholic steatohepatitis, even though Roux - en - Y gastric bypass was generally more effective at improving glycemic control, lipid profile, insulin resistance, and weight loss. This finding might be explained by the existence of a threshold in the weight loss or degree of metabolic improvement that is necessary to resolve nonalcoholic steatohepatitis.<sup>(54)</sup> In fact, the probability of reaching the primary endpoint increased non-linearly above 20% weight reduction and further decreases in bodyweight above this threshold translated into less additional histological improvement. Resolution of nonalcoholic steatohepatitis was also associated with postoperative improvement of insulin resistance and triglyceride concentrations.<sup>(54)</sup> A threshold mechanism for changes in insulin resistance might explain the lower effect of further weight reduction above 20% and the lack of differences observed between Roux - en - Y gastric bypass and sleeve gastrectomy.<sup>(54)</sup> A study investigating the effect of weight loss achieved through diet and physical exercise on liver histological features was done in 293 people with nonalcoholic steatohepatitis.<sup>(55)</sup> nonalcoholic steatohepatitis resolution was reached in 25% of the participants and 19% had regression of liver fibrosis.<sup>(55)</sup> The 1 - year mean weight loss in the study was 4.6 kg (SD 3.2); however, only 10% had a weight loss of 10% or greater.<sup>(55)</sup> Comparatively, the mean weight loss in the lifestyle - intervention group of this study was 5.5% and only 27% achieved a weight reduction of at least 10%. These findings provide reassurance regarding the effectiveness of lifestyle modification in our study, thus providing an appropriate comparator for the related effectiveness of surgical therapy. Studies show that new anti - obesity medications (eg, tirzepatide or cagrilintide plus semaglutide)<sup>(56,57)</sup> can achieve levels of weight loss close to 20% in some people, suggesting that these drugs might be more effective as a treatment of nonalcoholic steatohepatitis compared with lifestyle modification, as well as with pioglitazone and liraglutide, the drugs used in our study. Semaglutide achieved resolution of nonalcoholic steatohepatitis without fibrosis worsening in 59% of participants versus 17% in the placebo group in one trial. Importantly, however, there was no difference in previous studies between semaglutide and placebo in the downstaging of liver fibrosis,<sup>(58)</sup> despite the substantial weight loss achieved by this drug. This observation suggests that the net improvement of fibrosis achieved by surgery in our study might not be extrapolated to other forms of weight - loss interventions. The relative efficacy of newer anti - obesity

drugs on nonalcoholic steatohepatitis and liver fibrosis will therefore require further investigation.<sup>(59)</sup> The Intention - to - treat analysis showed that in participants with a nonalcoholic fatty liver disease activity score of 4 or 5 or more, the probability of reaching the primary endpoint was 3–5 times higher with bariatric - metabolic surgery than with lifestyle modification. In this subgroup, the improvement of at least one stage of fibrosis in the per protocol analysis was almost double after both Roux - en - Y gastric bypass and sleeve gastrectomy than after lifestyle modification. The ability of surgery to control and even improve fibrosis associated with nonalcoholic steatohepatitis is of particular clinical relevance given that fibrosis is the main predictor of liver complications and cardiovascular mortality and morbidity in nonalcoholic steatohepatitis.<sup>(59,60)</sup> The number of surgical complications in our study was similar after both surgical procedures. Several participants had gastroesophageal reflux after sleeve gastrectomy. Gastroesophageal reflux disease is a known complication of sleeve gastrectomy;<sup>(61)</sup> the high rate of gastroesophageal reflux disease in our study might partly be related to a more frequent use of postoperative diagnostic endoscopy compared with in usual clinical practice, which is a standard practice in postoperative assessment of patients undergoing sleeve gastrectomy at our centers. A cost - effectiveness analysis<sup>(62)</sup> of bariatric - metabolic surgery in individuals with nonalcoholic steatohepatitis showed that surgery is cost - effective in all individuals with obesity and nonalcoholic steatohepatitis, regardless of fibrosis stage, making surgery a suitable approach for the treatment of this condition. Previous studies had shown efficacy of bariatric - metabolic surgery on nonalcoholic fatty liver disease.<sup>(63 - 65)</sup> there is a study supports these findings and also provides evidence that benefits of surgery extend to nonalcoholic steatohepatitis and liver fibrosis. The results have important implications for clinical practice. There are no existing mechanisms for prioritization of bariatric - metabolic surgery in most health - care systems and access to surgery is often based on a first - come - first - served basis.<sup>(66)</sup> Other study supports prioritization of surgery in nonalcoholic steatohepatitis, especially in the presence of a high risk of liver - related morbidity and mortality. A study of 30 000 individuals with nonalcoholic fatty liver disease and BMI of 40 kg/m<sup>2</sup> or more showed that bariatric surgery conferred a 49% lower risk of cardiovascular disease compared with non - surgical care.<sup>(67)</sup> Whether or not surgery could be used as a treatment of nonalcoholic steatohepatitis in patients who do not meet standard criteria for bariatric - metabolic surgery cannot be extrapolated from our study and warrants further and specific investigation. Our study is the first to compare three active treatments of nonalcoholic steatohepatitis and investigate the efficacy of bariatric - metabolic surgery in a randomized trial. Importantly, this study used preoperative and postoperative liver biopsy, which is the gold standard for assessment of nonalcoholic steatohepatitis related endpoints.<sup>(68)</sup> This study has several limitations. The FDA guidance<sup>(68)</sup> was published recommending the use of a nonalcoholic fatty liver disease activity score of at least 4 with at least 1 point each in inflammation and ballooning along with a Fibrosis staging scored (CRN) fibrosis score of 2–3 as essential inclusion criteria in nonalcoholic steatohepatitis trials. In the present study, we included people with a nonalcoholic fatty liver disease activity score of at least 3, and inflammation and ballooning scores in line with the most recent FDA guidance.

However, consistent with most nonalcoholic steatohepatitis trials published since 2021 that study included people with fibrosis stages<sup>(69 - 73)</sup> to investigate the effect of interventions according to the most recent FDA recommendations, we did a post - hoc analysis of the primary endpoint in participants with a nonalcoholic fatty liver disease activity score of 4 or at least 5. We also did subgroup analyses of results in participants with fibrosis stages 2 and 3, who accounted for more than 50% of all participants in our study.<sup>(54)</sup> In the aggregate, the results of these subgroup analyses show that differences between surgical and non - surgical treatment of nonalcoholic steatohepatitis are greater among participants with more severe fibrosis. This finding supports the robustness of the overall findings of our study and their clinical relevance in patients with more advanced stages of fibrosis. As the study did not control for baseline BMI and glycemic control, differences in bodyweight and diabetes severity at baseline could have, at least partly, influenced the response to treatment.<sup>(54)</sup> However, BMI and HbA1c levels were higher in the Roux - en - Y gastric bypass group, which would potentially bias results against rather than in favor of surgery, which was the most effective intervention. Another limitation is that the medications used reflect indications and drugs available in Italy for people with obesity and nonalcoholic steatohepatitis at the time the study was designed.<sup>(54)</sup> Novel antiobesity drugs might result in better nonalcoholic steatohepatitis outcomes than those we observed in the non - surgical group of our study, given their greater weight - loss potential. Future research should compare new anti - obesity drugs with other active drugs or bariatric - metabolic surgery. Another important limitation is that all participants in this study were White, meaning that the rates of nonalcoholic steatohepatitis resolution and other metabolic improvements observed in this trial might not be generalizable to other ethnic groups.<sup>(54)</sup>

#### **Post - metabolic bariatric surgery associated with fewer cardiovascular mortality outcomes**

Obesity is associated with an increase incidence of heart failure, myocardial infarction, stroke, and death.<sup>(74 - 76)</sup> Weight loss has become a standard recommendation for all patients with cardiovascular disease and coexisting obesity; however, randomized evidence to support this recommendation is limited. The largest trial to date investigating whether intentional weight loss reduces cardiovascular morbidity and mortality in overweight individuals with type 2 diabetes did not show a significant reduction in cardiovascular events.<sup>(77)</sup> However, the weight loss observed in this trial was modest, and it is not known whether greater levels of weight loss would lead to clear reductions in recurrent cardiovascular events and deaths. Several studies of overweight and obese patients with cardiovascular disease suggest an “obesity paradox,” whereby elevated body mass index may be associated with lower mortality and cardiovascular events.<sup>(78, 79)</sup> However, other studies have described a U - shaped relationship with respect to weight, with severe obesity being associated with increased risk of cardiovascular outcomes.<sup>(76)</sup> Although the relationship between obesity and cardiovascular outcomes is not well delineated, bariatric surgery, in addition to being the most effective weight loss treatment, is also known to resolve other cardiovascular risk factors, including the burden of recurrent cardiovascular disease in patients with obesity and previous heart disease.<sup>(80)</sup> Randomized clinical

trials have revealed that bariatric surgery leads to consistent improvements in diabetes outcomes and cardiovascular risk factors in patients with diabetes and severe obesity.<sup>(81 - 83)</sup> However, such trials did not evaluate the effect of bariatric surgery on mortality and other major cardiovascular outcomes. Several observational studies have reported on the promising correlation between bariatric surgery and decreased cardiovascular outcomes.<sup>(32 - 85)</sup> Early studies of patients with severe obesity and type 2 diabetes undergoing bariatric surgery showed lower risks of macrovascular events in the surgery group, but confounding bias may have contributed to these results, and patients in these studies received procedures that are no longer common.<sup>(32)</sup> More recently, retrospective studies of patients with diabetes undergoing sleeve gastrectomy and Roux - en - Y gastric bypass have associated bariatric surgery with significantly lower incidence of incident major cardiovascular events after 8 years.<sup>(84, 85)</sup> More than 10 small randomized clinical trials have shown that metabolic surgery is superior to usual medical therapy for diabetes control and modifying cardiometabolic risk factors in patients with type 2 diabetes mellitus and obesity.<sup>(81 - 83, 86, 87)</sup> Furthermore, >30 large comparative cohort studies have consistently reported reduction in risk of mortality after metabolic surgery.<sup>(84, 85, 88)</sup> The majority of these Randomized clinical trials and large observational studies have only examined the favorable effects of Roux - en - Y gastric bypass (RYGB). Currently, sleeve gastrectomy (SG), a relatively new procedure, is the most commonly performed metabolic surgical procedure worldwide.<sup>(89 - 91)</sup> However, long - term data on efficacy of SG for macro - and microvascular complications of type 2 diabetes and mortality are limited.<sup>(92)</sup> Although cardiovascular and survival benefits of RYGB have been reported, the current study is among the first in the literature to show lower risk of coronary artery events, cerebrovascular events, heart failure, atrial fibrillation, nephropathy, and all case of mortality and mortality after SG compared with usual care.<sup>(92)</sup> The findings of this large retrospective study also provide evidence suggesting that RYGB in patients with obesity and type 2 diabetes may be associated with greater weight loss, better diabetes control, and lower risk of six - component coronary artery events, cerebrovascular events, heart failure, atrial fibrillation, nephropathy, and all case of mortality and mortality and nephropathy compared with SG. However, given the nature of the study, these data should be considered hypothesis generating and not conclusive.<sup>(92)</sup> Aristithes et al cohort study, demonstrated that bariatric surgery was associated with a significant lower incidence of coronary artery events, cerebrovascular events, heart failure, atrial fibrillation, nephropathy, and all case of mortality and mortality in patients with cardiovascular disease and severe obesity. These observed results apply to both patients with ischemic heart disease and heart failure. This study provides novel information and requires confirmation by a large scale randomize control trial, because effects of observational studies can be overestimated.<sup>(92)</sup>

#### **Post - metabolic bariatric surgery and decrease incidence of cancer and mortality**

Obesity and type 2 diabetes are both associated with increased cancer risk and cancer mortality.<sup>(94 - 99)</sup> In a study by Pearson - Stuttard et al.<sup>(99)</sup> an estimated 6% of all cancer cases worldwide in 2012 were attributable to the combined effects

of diabetes and overweight/ obesity. Additionally, the combination of diabetes and overweight/obesity contributed to as much as 25–40% of all liver cancer, esophageal adenocarcinoma, and endometrial cancer cases. Thus, the global pattern of increasing obesity and diabetes has led to an increased cancer prevalence. The causal link between obesity and 13 specific cancer types, referred to as obesity related, is well established.<sup>(100)</sup> In addition, some of these cancer types (i. e., liver, pancreatic, endometrial, colon and rectal, breast, bladder) have been suggested to be diabetes (mainly type 2) related.<sup>(101)</sup> However, at present, there is no clear consensus regarding which cancers are directly linked to type 2 diabetes.<sup>(102, 103)</sup> Obesity is a modifiable risk factor for type 2 diabetes and cancer, and bariatric surgery is the most effective intervention for substantial and sustained weight loss in patients with obesity.<sup>(104)</sup> Furthermore, the beneficial effects of bariatric surgery on type 2 diabetes are well described<sup>(81, 82)</sup> and the majority of patients with diabetes achieve short - term diabetes remission after bariatric surgery. Numerous observational and retrospective cohort studies of patients undergoing bariatric surgery have contributed to the understanding that intentional weight loss leads to reduced cancer risk and mortality in patients with obesity.<sup>(105 - 109)</sup> In a study by Ali Aminian et al, among adults with obesity, bariatric surgery compared with no surgery was associated with a significantly lower incidence of obesity - associated cancer and cancer - related mortality.<sup>(110)</sup> In conclusion, with increasing rates of obesity and diabetes worldwide, a greater emphasis on cancer prevention strategies is needed. Our results suggest that bariatric surgery may greatly reduce the risk of cancer among patients with obesity and diabetes. Moreover, durable diabetes remission seems imperative for cancer prevention in patients with obesity and diabetes.<sup>(111)</sup>

#### **Post - metabolic bariatric surgery increases life expectancy**

Lower mortality in association with bariatric surgery has also been observed in retrospective cohort studies, and a recent meta - analysis indicated 41% lower all - cause long - term  $\geq 2$  years mortality.<sup>(112)</sup> However, a recent retrospective study indicated that the relative risk of death remained higher among patients who underwent bariatric surgery than in the general background population.<sup>(113)</sup> Despite the beneficial effects of bariatric surgery, only a minority of eligible patients undergo such surgery.<sup>(114)</sup> It is recommended that patients considering bariatric surgery be given appropriate information to make an informed choice.<sup>(115)</sup> A major motivation to undergo bariatric surgery is to reduce the risks of deteriorating health and premature death.<sup>(116)</sup> However, the extent to which the reduced relative risk of death after bariatric surgery increases the life span in patients with obesity is unclear.<sup>(117)</sup> In 2007, our group provided prospectively collected data showing that mortality in a group of patients who underwent bariatric surgery was lower than in a matched control group of patients who were given usual care.<sup>(117)</sup> After an additional 13 years of follow - up, we now report that the difference in mortality persisted and that bariatric surgery was associated with a lower risk of death from both cardiovascular diseases and cancer. In addition, we show that life expectancy among patients with obesity who were given usual care was approximately 8 years shorter than in the general population, and this difference was decreased by bariatric surgery, which was associated with life

expectancy that was approximately 3 years longer than that associated with usual care. The mean increase in life expectancy after an intervention used for prevention of an outcome is generally lowest in populations at average risk, higher in populations with an elevated risk, and greatest among persons with established disease.<sup>(118)</sup> When viewed in the context of life - years gained with a variety of interventions in populations at elevated risk,<sup>(118)</sup> the 3 - year mean prolongation of life after bariatric surgery in our study is large. However, the controlled Swedish Obese Subjects study at baseline includes persons who have an elevated risk of serious and potentially life - threatening diseases, as well as persons with established conditions, such as type 2 diabetes and hypertension.<sup>(117)</sup> Therefore, it is important to emphasize that the reported gain in life expectancy is for our entire study population and cannot be translated into a survival benefit for a particular patient or for other populations beyond the study participants. The 8 - year - shorter life expectancy in the control group than in the general Swedish population is within the range previously reported for persons with severe obesity relative to those of healthy weight.<sup>(119 - 121)</sup> Although life expectancy was longer in association with bariatric surgery, survival was still lower than in the general population. This finding is consonant with a recent report from the Nordic countries showing that overall mortality among patients who had undergone bariatric surgery remained higher than in the general population.<sup>(122)</sup> Possible explanations for the remaining higher mortality in the surgery group include the above - normal BMI even after bariatric surgery, irreversible effects of obesity - related metabolic dysfunction that may have begun processes that lead to atherosclerosis or microvascular disease, and surgical complications and other factors causing death (alcoholism, suicide, and trauma). More frequent occurrence of alcoholism, suicide, and trauma<sup>(123 - 125)</sup> and a larger number of related deaths have previously been observed among patients with obesity who undergo bariatric surgery than among those who do not undergo surgery.<sup>(126)</sup> In the controlled Swedish Obese Subjects study, we have found a higher risk of alcohol abuse, suicide and self - harm, and serious fall - related injuries among patients with obesity who have undergone bariatric surgery than among those who have not.<sup>(127 - 130)</sup> In conclusion bariatric surgery was associated with lower overall mortality than usual obesity care. In middle - aged patients with severe obesity, life expectancy was approximately 3 years longer among patients who underwent surgery than among those who received usual care.<sup>(131)</sup>

#### **Post - Bariatric Weight Recurrence and Pharmacotherapy for Regain**

Bariatric surgery is a powerful tool for weight management and treating obesity - related comorbidities.<sup>(131 - 133)</sup> Yet weight regain occurs frequently a few years after bariatric surgery and it is often associated with recurrence of metabolic complications.<sup>(134, 135)</sup> Rates of weight regain vary across studies owing, in part, to variability in the definition.<sup>(135)</sup> One study reported that people regained an average of 22.5% of their lost weight at 3 years after surgery and more than two - thirds of them had regained at least 20% of their lost weight by 5 years.<sup>(135)</sup> Data from the Swedish Obesity Study suggest that weight loss is greatest at 1 to 2 years after bariatric surgery and rates of weight regain level off after 8 to 10 years. Currently, there is not a standardized or consistently utilized definition for post - bariatric weight regain.<sup>(136)</sup>

### 3. Conclusion

Obesity is a growing major health problem in populations. The lifestyle modification and pharmacotherapy remain important but are limited in weight loss results. Bariatric surgery is safe, effective and underutilized for treating obesity and related complication. Weight loss more than ten percent is often recommended. Metabolic bariatric surgery can effectively treat disease like type 2 diabetes and nonalcoholic fatty liver disease, and greater weight loss leads to greater remission in some group. In addition to atherosclerotic and cardiovascular risk reduction. Metabolic bariatric surgery may decrease cancer risk and increase life expectancy for people with obesity. promotes weight loss by modifying the gastrointestinal tract to restrict food intake, induce malabsorption. Bariatric surgery is associated with numerous complications, including the malabsorption of dietary of vitamins and minerals. Malabsorption can be averted through nutritional supplementation and periodic monitoring for deficiencies. Many complications can occur without sustained supplementation, of vitamins and minerals that induced by malabsorption bariatric surgery

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