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# The Efficacy of Roux-en-Y Gastric Bypass in Young-Onset Diabetes Mellitus: A Narrative Review

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### Abstract

Type 2 diabetes mellitus (T2DM) in youth is invading the communities because, if not controlled on time, the long-term complications include cardiovascular diseases, nephropathy, neuropathy, and retinopathy that cause immense mortality and morbidity. Lifestyle changes and antidiabetic drugs are considered the foundation of T2DM therapy. However, these adjustments usually do not effectively produce long-term glycemic regulation, especially in patients with obesity of the third and fourth degrees. Bariatric surgery has also been identified as an efficacious intervention for obesity and obesity-related complications such as T2DM. Roux-en-Y gastric bypass (RYGB) has proven to be one of the most effective procedures in causing considerable weight loss and enhancing glycemic changes. This review provides a comprehensive analysis of RYGB in patients with young-onset T2DM regarding the improvement of glycemic control, weight loss, and diabetes comorbidities. RYGB has been established as a practice in the treatment of T2DM and severe obesity. This narrative review underscores the various effects of RYGB, such as enhanced glycemic control, considerable and long-term weight loss, and reduced cardiovascular disease risks. However, the review also points toward the directions and the adverse effects of RYGB regarding metabolic and skeletal health. There are risks of nutritional deficiencies, increased fracture rates, and even relapse to diabetes, which make patient selection, proper pre and postoperative investigation, and critical monitoring.

Categories: Anesthesiology

Keywords: bariatric surgery, glycemic control, weight loss, roux-en-y gastric bypass, type 2 diabetes mellitus

## **Introduction And Background**

Type 2 diabetes mellitus (T2DM) is one of the most common chronic diseases, characterized by insulin resistance and insufficient insulin production by the beta-cells of the pancreas that result in elevated blood glucose levels. It is widely considered a significant health problem, which is prevalent and on the rise, especially among the youth. T2DM in youth is invading the communities because, if not controlled on time, the long-term complications include cardiovascular diseases, nephropathy, neuropathy, and retinopathy that cause immense mortality and morbidity [1-4]. Lifestyle changes and antidiabetic drugs are considered the foundation of T2DM therapy. However, these adjustments usually do not effectively produce long-term glycemic regulation, especially in patients with third and fourth-degree obesity. Bariatric surgery has been identified as an efficacious intervention for obesity and obesity-related complications such as T2DM. Of the different bariatric operations, Roux-en-Y gastric bypass (RYGB) has proven to be one of the most effective procedures in causing considerable weight loss and enhancing glycemic changes [5,6].

RYGB is a surgical procedure wherein the surgeon develops a small pouch in the stomach, and a section of the small intestine is bypassed. This change of behavior not only limits the consumption of food but directs hormonal changes that promote the effectiveness and volume of insulin and secretion, yielding almost immediate and effective management of glycemia. The effectiveness of RYGB surgery in the treatment of or contribution to the remission or significant amelioration of overweight/obesity-associated T2DM in the adult population is abundantly clear in the literature; however, its implication for young-onset T2DM remains relatively unknown [7,8]. Numerous global medical and surgical organizations have agreed that bariatric surgery should be included in the treatment options for type 2 diabetes in adults, as it has been used for a long time as a therapy for severe obesity. Vertical sleeve gastrectomy (VSG), adjustable gastric banding (AGB), and RYGB are the most often used bariatric procedures in both adults and adolescents. RYGB and VSG are the most successful among them in terms of improving metabolic health and lowering body weight [9].

This review provides a comprehensive analysis of RYGB in patients with young-onset T2DM concerning the improvement of glycemic control, weight loss, and diabetes comorbidities. This review aims to reveal the advantages and possible drawbacks of RYGB as a treatment for youths with T2DM based on the accumulated research findings.





### Review

The efficacy of RYGB for managing T2DM has been extensively studied, with numerous studies demonstrating its benefits. Giudicelli et al. found that RYGB enhances metabolic health and diabetes outcomes in both young and older non-insulin-dependent diabetic adults, emphasizing the importance of timely surgery for better glycemic control and reduced cardiovascular risk [10]. Hage et al. reported that RYGB is more effective than sleeve gastrectomy (SG) for T2DM remission, regardless of baseline characteristics and severity [11].

Girundi et al. concluded that RYGB is highly effective for long-term glycemic control, with 87.6% of patients experiencing T2DM remission 18 months post-surgery [12]. Cheng et al. demonstrated that RYGB is more effective than medical treatment for achieving glycemic control over five years, with a higher probability of maintaining HbA1c levels below 6.5% without medication [13]. Ji et al. showed that RYGB benefits T2DM patients with a body mass index (BMI) under 32.5 kg/m<sup>2</sup>, improving weight loss, glycemic control, and lipid profiles over a six-year follow-up [14].

Nudotor et al. found that RYGB is significantly more effective than VSG for sustained T2DM remission over five years [15]. van Rijswijk et al. reported that both RYGB and one-anastomosis gastric bypass are effective for rapid T2DM remission in severely obese patients [16]. Du et al. confirmed the safety, efficacy, and practicality of laparoscopic RYGB for treating obese Chinese T2DM patients over a three-year follow-up [17]. Zhang et al. found that RYGB is superior to SG for medication decrease and dyslipidemia remission [18].

Sahin et al. highlighted the need to prioritize patient outcomes over economic considerations in non-obese overweight groups undergoing bariatric surgery. Collectively, these studies indicate that RYGB is highly effective for inducing T2DM remission, improving glycemic control, and achieving significant weight loss, outperforming other bariatric procedures such as SG and VSG in most metrics, making it a valuable option for managing T2DM in various patient populations [19]. van de Pas et al. conducted a nationwide cohort study on 2,822 young adults and 24,497 adults, revealing that bariatric surgery, specifically RYGB or SG, is equally safe and effective across age groups, with young adults achieving superior weight loss. Guimarães et al. found that RYGB effectively promotes long-term weight loss and improves obesity-related comorbidities with minimal serious complications in a study of 601 patients. Sharples et al. demonstrated that RYGB significantly improved diabetic control, reduced medication use, and achieved substantial weight loss and cost savings in 52 patients with T2DM [20-22].

#### Advantages of Roux-en-Y gastric bypass

RYGB offers significant advantages in T2DM by inducing improvements in glucose metabolism and insulin sensitivity, as well as enhancing beta-cell function. Research indicates that RYGB leads to early increases in insulin secretion and sustained improvements in glucose tolerance and insulin sensitivity, which are maintained even seven years post-surgery [23-25]. Furthermore, RYGB triggers changes in beta-cell mass, promoting trans-differentiation into alpha-cell mass due to long-term adaptation of the digestive tract post-surgery, contributing to the resolution of T2DM. Additionally, the effects of the procedure on s-cell function are crucial for the long-term management of T2DM, highlighting the lasting benefits of RYGB in treating this metabolic disorder [26]. RYGB specifically improves insulin sensitivity in the liver, muscle, and fat, as evidenced by increased glucose disposal and decreased endogenous glucose production [27]. Furthermore, RYGB induces marked improvements in glucose tolerance, insulin sensitivity in muscle and liver, and beta-cell function, which are maintained even up to seven years post-surgery [28].

RYGB significantly impacts ghrelin levels, a hormone associated with hunger regulation. Research indicates that both RYGB and laparoscopic sleeve gastrectomy (LSG) lead to notable reductions in serum ghrelin concentrations one-year post-surgery, with decreases of approximately 0.068 pg/mL for RYGB and 0.083 pg/mL for LSG. However, these changes were not statistically different between the two procedures. Additionally, a study on mice demonstrated that while ghrelin levels increased post-surgery, lower ghrelin levels did not affect the metabolic benefits of RYGB, suggesting that other mechanisms may also contribute to weight loss and metabolic improvements. Furthermore, ghrelin modulation has been explored as a potential therapeutic strategy to enhance weight loss outcomes in patients with suboptimal responses to bariatric surgery. Overall, RYGB effectively reduces ghrelin levels, contributing to its role in weight management and metabolic health [29,30].

#### Disadvantages of Roux-en-Y gastric bypass

Despite initial improvements, many patients relapse within five years due to changes in the beta-cell population and absorptive intestinal effects. This leads to beta-cell exhaustion and a decrease in beta-cell mass. Risks associated with RYGB include higher rates of mortality and morbidity compared to other bariatric procedures, long-term nutritional deficiencies (such as bone demineralization), and potential complications such as ulcers, bowel obstructions, and hypoglycemia syndromes [31-33]. Dumping syndrome is a common issue post-RYGB, characterized by symptoms such as borborygmus, diarrhea, abdominal distention, pain, palpitations, tachycardia, hypotension, sweating, flushing, and fatigue, typically occurring within 30 minutes of eating. A Danish questionnaire-based study by Grabsholt et al. found that 52.4% of



1,429 participants who had RYGB 4.7 years prior reported experiencing one or more dumping syndromerelated symptoms [33,34].

Nutritional deficiencies and altered nutrient absorption are significant concerns after RYGB, necessitating lifelong supplementation of vitamin B12, vitamin D, calcium, iron, and folic acid. Anemia, affecting about 12% of patients at baseline, is the most prevalent clinical symptom of these deficiencies. Additionally, there is concern that RYGB impairs bone metabolism, increasing the risk of fractures, osteoporosis, and osteopenia [35,36]. Table *1* depicts a description of the studies included in the review.

| Author<br>and year                  | Participants  | Intervention  | Outcome measures  | Conclusion  |
|-------------------------------------|---|---|---|---|
| Giudicelli<br>et al.<br>(2023) [10] | 169 non-<br>insulin-<br>dependent<br>diabetic<br>adults who<br>underwent<br>RYGB. 55<br>young adults<br>and 114<br>older adults | To ensure group comparability,<br>young adults were matched one to<br>two with older adults based on<br>sex, BMI, duration of preoperative<br>diabetes, and the American<br>Society of Anesthesiologists score                            | Primary: Diabetes<br>remission, focusing on<br>rates and timing of<br>remission post-surgery.<br>Secondary: Percent weight<br>change, loss to follow-up,<br>and all-cause adverse<br>events                               | The results support the efficacy of RYGB<br>in enhancing metabolic health and<br>diabetes outcomes in both young and older<br>non-insulin-dependent diabetic adults,<br>emphasizing the importance of timely<br>metabolic surgery for better glycemic<br>control and reduced cardiovascular risk  |
| Hage et al.<br>(2024) [11]          | 1,149<br>patients with<br>T2DM who<br>underwent<br>either RYGB<br>or SG   | The study evaluated the rates of<br>T2DM remission, insulin use,<br>changes in HbA1c levels, and<br>diabetes medication usage for<br>both RYGB and SG patients to<br>determine the effectiveness of<br>each procedure in managing<br>T2DM | Diabetes remission<br>comparison, weight change,<br>and adverse events and<br>long-term metabolic<br>outcomes   | RYGB has higher efficacy for T2DM<br>remission than SG, regardless of baseline<br>characteristics, T2DM severity, weight<br>loss, or follow-up duration   |
| Girundi<br>(2016) [12]              | 468 patients<br>with T2DM<br>who<br>underwent<br>RYGB   | Patients' glycemic profiles were<br>assessed at specific intervals after<br>surgery, including the 3rd, 6th,<br>9th, 12th, and 18th months  | Remission of T2DM   | The study concluded that RYGB was<br>highly effective in achieving long-term<br>glycemic control, with 87.6% of patients<br>experiencing remission of type 2 diabetes<br>18 months after the surgery  |
| Cheng et<br>al. (2022)<br>[13]      | 100 adults<br>with T2DM   | Throughout the five-year study,<br>participants were periodically<br>examined to evaluate the results<br>and monitor for any side effects   | HbA1c levels of less than<br>6.5%, weight loss, changes<br>in cardiovascular risk<br>factors, quality of life, and<br>adverse events  | Over 5 years, in adults with type 2<br>diabetes mellitus and a BMI between 27<br>and 32 kg/m <sup>2</sup> , RYGB was more effective<br>than the medical treatment for achieving<br>glycemic control. Individuals undergoing<br>RYGB surgery had a higher probability of<br>attaining HbA1c levels below 6.5% without<br>requiring diabetes medication in<br>comparison to those undergoing medical<br>treatment |
| Ji et al.<br>(2020) [14]            | 52 patients<br>with T2DM  | Data were collected at baseline, at<br>3, 6, and 1, 2, 3, 4, 5, and 6 years<br>after surgery to monitor changes<br>in the measured parameters over<br>time  | Throughout the 6-year<br>study period, alterations in<br>lipid profiles (triglycerides,<br>total cholesterol, LDL, HDL,<br>BMI, HbA1c levels, OGTT<br>results, and the remission<br>or improvement of T2DM)<br>were noted | Results indicate that T2DM patients with a<br>BMI < 32.5 kg/m <sup>2</sup> might benefit from<br>RYGB, which can result in notable weight<br>loss, better glycemic control, and positive<br>alterations in lipid profiles   |
| Nudotor et<br>al. (2021)<br>[15]    | 1,364<br>patients with<br>T2DM  | The study compared the efficacy<br>of VSG and RYGB for long-term<br>T2DM remission. The study used<br>a retrospective cohort study<br>design  | T2DM remission, HbA1c<br>less than 6.5%   | Over a five-year follow-up period, it was<br>discovered that RYGB was significantly<br>more effective than VSG in achieving<br>sustained remission of T2DM  |
| Van                                 | 220 severely  | A randomized controlled clinical<br>trial was undertaken to assess the<br>comparative efficacy of two   | Glycemic control at 12<br>months follow-up, weight  | Both RYGB and OAGB were effective in  |





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| Rijswijk et<br>al. (2022)<br>[16]   | obese<br>patients with<br>T2DM  | distinct bariatric surgical<br>procedures: laparoscopic RYGB<br>and laparoscopic OAGB,<br>specifically concerning glycemic<br>control  | loss, surgical complications,<br>psychologic status, and<br>quality of life  | inducing rapid remission of T2DM in<br>severely obese patients, as highlighted in<br>the trial protocol   |
|-------------------------------------|---|--|--|---|
| Du et al.<br>(2018) [17]            | 58 class 1<br>obese<br>patients with<br>T2DM                                    | Laparoscopic RYGB surgery, a<br>minimally invasive weight loss<br>and diabetes management<br>treatment, was performed on all<br>patients. The efficiency of<br>laparoscopic RYGB in causing<br>weight loss and T2DM remission<br>was evaluated using outcome<br>measures   | Waist circumference,<br>fasting plasma glucose<br>levels, and outcomes of an<br>OGTT at 2 hours  | Over a 3-year follow-up period,<br>laparoscopic RYGB was proven to be safe,<br>efficacious, and practical in treating obese<br>Chinese class I patients with T2DM   |
| Zhang et<br>al. (2020)<br>[18]      | 35<br>participants<br>who<br>underwent<br>RYGB and<br>70 who<br>underwent<br>SG | Participants who received RYGB<br>were matched with up to two SG<br>participants based on specified<br>factors such as age, sex, body<br>mass index, HbA1c level,<br>medication use, duration of<br>diabetes, and blood pressure. All<br>surgical procedures, including SG<br>and RYGB, were carried out<br>laparoscopically | Rate of diabetes medication<br>discontinuation, changes in<br>serum cholesterol and low-<br>density lipoprotein-c levels,<br>and diabetes control within<br>the 24-month follow-up<br>period   | Both RYGB and SG were great procedures<br>for treating obesity in patients with type 2<br>diabetes; however, RYGB was linked to<br>better outcomes than SG in terms of<br>medication decrease related to metabolic<br>disorders and dyslipidemia remission                  |
| Sahin et<br>al. (2018)<br>[19]      | 54 patients<br>with T2DM  | Participants who underwent the<br>laparoscopic RYGB procedure<br>had type 2 diabetes mellitus. The<br>BMI of these individuals was then<br>used to categorize them into<br>groups: above or below 30 kg/m <sup>2</sup>   | Changes in HbA1c levels,<br>weight loss progression,<br>and improvement in<br>diabetes symptoms  | The study questioned the suitability of<br>cost-effectiveness analysis for non-obese<br>overweight groups undergoing bariatric<br>surgery, highlighting the need to prioritize<br>patient outcomes and mortality reduction<br>over economic considerations in such<br>cases |
| van de pas<br>et al.<br>(2023) [20] | 2,822 young<br>adults and<br>24,497<br>adults who<br>underwent<br>RYGB or SG    | A nationwide population-based<br>cohort study utilizing data from the<br>Dutch Audit Treatment of Obesity.<br>Young adults (aged 18–25 years)<br>and adults (aged 35–55 years)<br>who underwent primary RYGB or<br>SG were included  | The primary outcome was<br>the percentage total weight<br>loss measured up to five<br>years postoperatively.<br>Secondary outcomes<br>included the improvement<br>of obesity-related<br>comorbidities such as<br>hypertension, dyslipidemia,<br>and musculoskeletal pain | Bariatric surgery was found to be equally<br>safe and effective in young adults as in<br>adults, based on the study's findings.<br>Young adults showed superior weight loss<br>outcomes compared to adults after<br>undergoing RYGB or SG procedures                        |
| Guimaraes<br>et al.<br>(2021) [21]  | 601 patients<br>who<br>underwent<br>RYGB  | The study evaluated the long-term<br>efficacy of RYGB in patients who<br>underwent the procedure for<br>obesity treatment  | Percentage total weight<br>loss, percentage excess<br>weight loss, percentage<br>excess BMI loss, rates of<br>comorbidity remission, and<br>complications associated<br>with RYGB  | RYGB is an effective long-term solution for<br>weight loss and the improvement of<br>obesity-related comorbidities, with a low<br>incidence of serious complications  |
| Sharples<br>et al.<br>(2019) [22]   | 52 patients with T2DM   | The study reviewed the records of<br>patients from a bariatric clinic who<br>had a confirmed diagnosis of type<br>2 diabetes mellitus and underwent<br>RYGB  | Weight loss, HbA1c levels,<br>medication use, and cost<br>analysis   | The study concluded that RYGB is<br>effective in improving diabetic control,<br>reducing medication use, facilitating<br>significant weight loss, and providing<br>substantial cost savings for obese patients<br>with type 2 diabetes                                      |

#### TABLE 1: Description of the studies included in the review.

RYGB: Roux-en-Y gastric bypass, T2DM: Type 2 Diabetes Mellitus, HbA1c: Glycated haemoglobin, BMI: Body Mass Index, LDL: Low-density lipoprotein, HDL: High-Density Lipoprotein, OGTT: Oral glucose tolerance test, VSG: vertical sleeve gastrectomy, OAGB: one-anastomosis gastric bypass, SG: Sleeve gastrectomy



#### Discussion

The studies reviewed offer a comprehensive evaluation of the effectiveness and safety of RYGB and SG in managing obesity and T2DM. van de Pas et al. (2023) conducted a large-scale study on young and older adults, demonstrating that bariatric surgery is equally safe and effective across age groups. Young adults achieved superior weight loss outcomes compared to adults, with significant improvements in obesity-related comorbidities such as hypertension, dyslipidemia, and musculoskeletal pain. This finding suggests that age should not be a limiting factor when considering candidates for bariatric surgery, as both young and older adults benefit substantially [20].

Comparative studies between RYGB and SG, such as those by Hage et al. (2024) and Zhang et al. (2020), highlight that RYGB generally offers higher efficacy for T2DM remission than SG. Hage et al. (2024) found that RYGB leads to better diabetes remission rates, reduced insulin use, and improved HbA1c levels compared to SG, regardless of patients' baseline characteristics. Zhang et al. (2020) supported these findings, reporting that RYGB was more effective in reducing medication use and improving metabolic parameters such as serum cholesterol and low-density lipoprotein levels. These studies underscore the superiority of RYGB over SG in achieving more favorable metabolic outcomes in T2DM patients [11,18]. Several studies focused on the long-term efficacy of RYGB in managing T2DM and obesity. Girundi (2016) found that 87.6% of patients experienced T2DM remission 18 months post-RYGB, indicating significant long-term glycemic control. Similarly, Cheng et al. (2022) showed that RYGB was more effective than medical treatment alone in achieving and maintaining HbA1c levels below 6.5% over five years. Ji et al. (2020) and Du et al. (2018) extended these findings by demonstrating sustained weight loss, improved glycemic control, and positive changes in lipid profiles over multi-year follow-up periods, further establishing the long-term benefits of [12-14].

Giudicelli et al. (2023) and van Rijswijk et al. (2022) highlighted the importance of timely metabolic surgery for improving glycemic control and reducing cardiovascular risks in both young and older adults with T2DM. Giudicelli et al. (2023) emphasized that RYGB enhances metabolic health and diabetes outcomes, while van Rijswijk et al. (2022) compared RYGB and one-anastomosis gastric bypass, concluding that both procedures were effective in inducing rapid T2DM remission in severely obese patients. These studies illustrate the critical role of bariatric surgery in managing severe obesity and its associated comorbidities, regardless of the specific surgical procedure used [10,15].

#### **Recommendations and future directions**

To strengthen the evidence base, future research should prioritize randomized controlled trials (RCTs) that compare the effectiveness of RYGB and SG. RCTs can reduce selection bias and provide more robust conclusions about the comparative benefits and risks of these procedures. Developing and implementing standardized outcome measures across studies will facilitate easier comparisons and a clearer synthesis of results. Consistent reporting on key metrics such as percentage total weight loss, glycemic control, remission rates of T2DM, and improvement in obesity-related comorbidities is essential. This standardization will improve the overall quality of meta-analyses and systematic reviews. Future studies should extend the follow-up period beyond five years to assess the long-term sustainability of weight loss and metabolic improvements. This will provide a more comprehensive understanding of the durability of the benefits and the potential for late-onset complications. Research should include detailed analyses of postoperative nutritional monitoring and patient adherence to supplementation regimens. Investigating strategies to improve compliance with vitamin and mineral supplementation will help mitigate the risk of nutritional deficiencies.

Investigating the factors that predict success with different bariatric procedures can lead to more personalized treatment approaches. Identifying patient characteristics, such as genetic, metabolic, and behavioral factors, that influence outcomes can help tailor surgical options to individual needs. Personalized treatment plans can enhance the efficacy and safety of bariatric surgery for each patient.

## Conclusions

RYGB has been established as a practice in the treatment of T2DM and severe obesity. This systematic review underscores the various effects of RYGB, such as enhanced glycemic control, considerable and long-term weight loss, and reduced cardiovascular disease risks. The data in the present and previous work imply that RYGB is more efficient in diabetes remission and better influences metabolic profiles compared to conventional and intensive medical therapies; such advantageous impacts are applicable in young and elderly populations as well. However, the review also points toward the directions and the adverse effects of RYGB regarding metabolic and skeletal health. There are risks of nutritional deficiencies, increased fracture rates, and even diabetes relapse, which make patient selection, proper pre and postoperative investigation, and monitoring critical. Therefore, the lifelong necessity for the intake of supplements and the frequency of monitoring further complications relating to bone health are essential aspects to consider in patients who have undergone RYGB.

Further studies should be conducted on patient selection criteria to maximize the benefits of RYGB, post-





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surgery supportive care, and the genetic and molecular basis of the variability of RYGB outcomes. If all these challenges are managed, then the medical establishment will be able to optimize RYGB benefits concerning patients with T2DM and obesity. Therefore, RYGB is an effective treatment in the fight against T2DM and obesity, with the outcomes demonstrating improvements in metabolic and cardiovascular profiles. However, strategies that look at the advantages as well as drawbacks of various procedures are crucial for achieving the best outcomes for the patients.

### **Additional Information**

#### **Author Contributions**

All authors have reviewed the final version to be published and agreed to be accountable for all aspects of the work.

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