WJD

# World Journal of **Diabetes**

Submit a Manuscript: https://www.f6publishing.com

World J Diabetes 2024 November 15; 15(11): 2157-2161

DOI: 10.4239/wjd.v15.i11.2157

ISSN 1948-9358 (online)

EDITORIAL

## Diabetes and obesity: A debate on bariatric interventions and its implications

#### José C Tatmatsu-Rocha, Marcos R Lima da Silva

Specialty type: Endocrinology and metabolism

Provenance and peer review: Invited article; Externally peerreviewed.

Peer-review model: Single-blind

Peer-review report's classification Scientific Quality: Grade B, Grade D, Grade D Novelty: Grade B, Grade C, Grade С Creativity or Innovation: Grade B, Grade C, Grade C Scientific Significance: Grade B, Grade C, Grade C

P-Reviewer: Bedi DS; De Zoysa W; Sachan A

Received: April 3, 2024 Revised: August 29, 2024 Accepted: September 26, 2024 Published online: November 15, 2024 Processing time: 195 Days and 20.6

Hours

José C Tatmatsu-Rocha, Marcos R Lima da Silva, College of Medicine, Postgraduate Program in Physiotherapy and Functionality, Department of Physiotherapy, Federal University of Ceará-UFC, Fortaleza 60430-450, Ceará, Brazil

Corresponding author: José C Tatmatsu-Rocha, MSc, PhD, Professor, Research Scientist, College of Medicine, Postgraduate Program in Physiotherapy and Functionality, Department of Physiotherapy, Federal University of Ceará-UFC, Rua Major Weyne, 1440, Fortaleza 60430-450, Ceará, Brazil. tatmatsu@ufc.br

## Abstract

In this editorial, we comment on an article by Tang et al published in the World *Journal of Diabetes.* Obesity and diabetes are two pathological situations that are intrinsically related. Neither lifestyle changes nor pharmacological treatments have achieved diabetes remission. From this perspective, bariatric surgery has been widely used as an approach for weight loss in obese patients and as a strategy to promote metabolic modulation. The main effects of bariatric surgery involve direct action in improving cardiovascular function and endothelial function and reducing insulin resistance, leading to diabetes remission in the short term following surgery. In this context, it has been observed that hormones from the gastrointestinal tract and endothelium play a prominent role in this process. By reversing endothelial dysfunction, it is possible to balance pro-inflammatory cytokine production, improving the availability of nitric oxide and inhibiting vascular oxidative stress. Furthermore, it can be considered an efficient anti-inflammatory strategy, alleviating interferon-gamma-mediated adipose tissue inflammation. The current challenge must be to unravel the pathophysiological mechanisms and potential targets for treating metabolic diseases.

Key Words: Bariatric surgery; Diabetes mellitus; Oxidative stress; Endothelial dysfunction; Obesity

©The Author(s) 2024. Published by Baishideng Publishing Group Inc. All rights reserved.



WJD | https://www.wjgnet.com

**Core Tip:** In this editorial, we address the pathophysiological mechanisms associated with bariatric surgery in type 2 diabetes remission. Within this perspective, we discuss gastrointestinal tract hormones, mainly peptide-1, which is a hormone secreted by gastrointestinal L cells and released immediately after food intake, as well as the benefits obtained after reversing endothelial syndrome *via* bariatric surgery.

**Citation:** Tatmatsu-Rocha JC, Lima da Silva MR. Diabetes and obesity: A debate on bariatric interventions and its implications. *World J Diabetes* 2024; 15(11): 2157-2161 **URL:** https://www.wjgnet.com/1948-9358/full/v15/i11/2157.htm

**DOI:** https://dx.doi.org/10.4239/wjd.v15.i11.2157

#### INTRODUCTION

Obesity is a disease characterized by excessive accumulation of body fat, and it has been associated with adverse metabolic conditions[1]. It directly contributes to increased cardiovascular risk with increased mortality[2]. Among the main changes caused by obesity are insulin resistance, arterial hypertension, dyslipidemia and chronic inflammation[3, 4]. Diabetes and obesity have generated a great burden on the health system, and today they are considered epidemic diseases<sup>[1]</sup> and have incapacitated some economically active individuals<sup>[5]</sup>. The impacts on the quality of life of this group must be taken into consideration, given that obesity and diabetes are health conditions generally diagnosed simultaneously[6]. The literature has shown that obesity promotes a significant increase in mortality related to metabolic diseases[7]. Insulin resistance, its high rates, decreased hormonal regulation and systemic inflammation are pathophysiological characteristics that make obesity and diabetes close realities<sup>[8]</sup>. Insulin resistance occurs due to the increased release of fatty acids, lipids and other advanced factors from adipose tissue, leading to numerous complications [9]. In this context, the need for disease control is clear, as diabetes management goes far beyond glycemic control. Maintaining an ideal weight is essential to prevent or reduce the incidence of macro and microvascular diseases and their complications[2,10]. Even in the face of the constantly developing pharmaceutical industry, with numerous promising drugs, studies point to bariatric surgery as a promising and potentially curative approach for the treatment of obesity and type 2 diabetes mellitus[11]. Also described as metabolic surgery, bariatric surgery has shown efficacy in providing prolonged remission of diabetes without the need for continuous pharmacological treatment[12]. Among patients undergoing bariatric surgery, approximately 75%-80% of patients showed significant improvement in type 2 diabetes control at 1-year follow-up appointments[12,13]. The proportion of patients who achieved diabetes remission across studies was estimated to be between 30 and 77% [13]. However, this proportion decreased when a longer follow-up period was performed in patients with chronic diabetes[14]. There are validated tools that have been used preoperatively to predict whether the patient will have diabetes remission after bariatric surgery. These include the DiaRem score, which has predictive power based on variables such as age, glycated hemoglobin values, insulin use, and use of other blood glucose- reducing agents[15].

#### BARIATRIC SURGERY: WHAT HAS CHANGED OVER TIME

Over the decades, bariatric surgery has proven to be very effective in achieving significant and sustainable weight loss in many patients[16]. Laparoscopic techniques have contributed to access to new techniques in bariatric surgery in the last decade. Some techniques have been transformative in the field. These include vertical banded gastroplasty[17], which decreased at the end of the 1990s and the adjustable laparoscopic gastric band[18], which emerged around 2012. From then on, biliopancreatic bypass[19] decreased compared to Roux-en-Y gastric bypass (RYGB)[20]. Finally, the use of laparoscopic sleeve gastrectomy[21] has rapidly increased in recent years.

#### MECHANISMS OF PHYSIOLOGICAL ADJUSTMENTS AFTER BARIATRIC SURGERY

The described mechanisms regarding the effects of bariatric interventions on glycemic index remission are still not clear. Most studies indicate that the weight loss provided by surgery is a determining factor in the resolution of type 2 diabetes [22]. Therefore, patients who experience greater weight loss are more likely to control or even remit the disease compared to those with less weight loss. Other variables include the patient's age, duration of illness and glycemic control, which are mechanisms independent of weight loss[4]. The main marker of diabetes resolution for years has been weight loss. However, there is a growing trend towards the use of metabolic markers, such as an increase in cardiovascular risk factors, endothelial function and lipid profile[23].

RYGB surgery has been increasingly used because it is known to be beneficial in weight loss and in reducing deaths from cardiovascular complications[24]. Glucagon-like peptide-1 (GLP-1) has been considered a key metabolic variable to verify clinical improvement after RYGB, whose effects (GLP-1) are independent of weight. This peptide has protective effects on the endothelium in addition to its metabolic actions. After RYGB surgery, there is a marked increase in GLP-1,

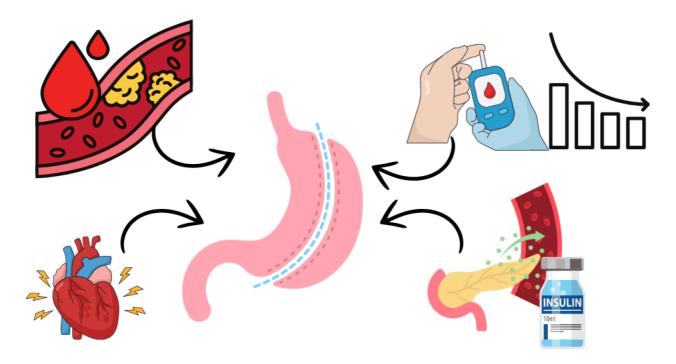


Figure 1 The effects of bariatric surgery involve a direct action on improving cardiovascular function, improving endothelial function and reducing insulin resistance, leading to diabetes remission in the short term after surgery.

which does not occur with weight loss due to dietary restriction. In this sense, a hypothesis has been formulated that GLP-1 is altered in the face of rapid metabolic changes and that this would only occur after RYGB, given that insulin resistance occurs even before weight loss after bariatric intervention [25,26].

## ENDOTHELIAL EFFECTS OF BARIATRIC SURGERIES

There are reports that the visceral adipose tissue of patients with obesity presents gene expression related to inflammation, oxidative stress and production of pro-inflammatory cytokines. This leads to impaired arteriolar function and possibly endothelial dysfunction[27], which can cause important cardiovascular changes. When stimulated, the endothelium can release agents that cause vasomotor function and hemostasis[27,28]. In contrast, the endothelium in healthy conditions presents low levels of oxidative stress and endothelial smooth muscle with relaxed tone due to the release of some markers that act on this vascular smooth muscle. This balance of vasodilators and vasoconstrictors as well as procoagulant and anticoagulant agents is essential for the ideal physiological balance of the endothelium. Endothelial dysfunction occurs due to an imbalance of these agents and evolves into some cardiovascular pathologies, such as cerebrovascular diseases, as well as peripheral arterial disease, coronary artery disease and type 2 diabetes[29,30].

## CONCLUSION

Bariatric surgeries may be beneficial in protecting against cardiovascular disease, especially when performed early, before the development of endothelial lesions (Figure 1). However, hormonal changes play a crucial role in diabetes remission. There is an urgent need for in-depth research that fully explains the metabolic and hormonal effects of the gastrointestinal tract and the different perspectives on the mechanisms underlying this approach.

## FOOTNOTES

Author contributions: Tatmatsu-Rocha JC and Lima da Silva MR contributed to writing and editing the manuscript and reviewing the literature; Tatmatsu-Rocha JC designed the overall concept and draft of the manuscript; Lima da Silva MR contributed to the discussion and conception of the manuscript; Lima da Silva MR revised the final version of the manuscript.

Conflict-of-interest statement: The authors declare that they have no conflict of interest.

**Open-Access:** This article is an open-access article that was selected by an in-house editor and fully peer-reviewed by external reviewers. It is distributed by the Creative Commons Attribution-NonCommercial (CC BY-NC 4.0) license, which permits others to distribute,



Zaishideng® WJD | https://www.wjgnet.com

remix, adapt, build upon this work non-commercially, and license their derivative works on different terms, provided the original work is properly cited and the use is non-commercial. See: https://creativecommons.org/Licenses/by-nc/4.0/

#### Country of origin: Brazil

ORCID number: José C Tatmatsu-Rocha 0000-0003-4900-4857; Marcos R Lima da Silva 0000-0003-3875-7044.

S-Editor: Fan M L-Editor: Filipodia P-Editor: Zheng XM

## REFERENCES

- 1 Yin M, Wang Y, Han M, Liang R, Li S, Wang G, Gang X. Mechanisms of bariatric surgery for weight loss and diabetes remission. J Diabetes 2023; 15: 736-752 [PMID: 37442561 DOI: 10.1111/1753-0407.13443]
- Tatmatsu-Rocha JC, Mendes-Costa LS. Inflammatory markers, oxidative stress, and mitochondrial dynamics: Repercussions on coronary 2 artery disease in diabetes. World J Diabetes 2024; 15: 1853-1857 [PMID: 39280176 DOI: 10.4239/wjd.v15.i9.1853]
- Powell-Wiley TM, Poirier P, Burke LE, Després JP, Gordon-Larsen P, Lavie CJ, Lear SA, Ndumele CE, Neeland IJ, Sanders P, St-Onge MP; 3 American Heart Association Council on Lifestyle and Cardiometabolic Health; Council on Cardiovascular and Stroke Nursing; Council on Clinical Cardiology; Council on Epidemiology and Prevention; and Stroke Council. Obesity and Cardiovascular Disease: A Scientific Statement From the American Heart Association. Circulation 2021; 143: e984-e1010 [PMID: 33882682 DOI: 10.1161/CIR.000000000000973]
- 4 Xu H. Obesity and metabolic inflammation. Drug Discov Today Dis Mech 2013; 10 [PMID: 24003334 DOI: 10.1016/j.ddmec.2013.03.006]
- Segateli L, Oliveira MAD, Coca MLL, Fogaça ECBSES, Silva LIB, Silva PL, Rocha Junior PR, Lopes LDG. Estratégias de promoção à saúde 5 para pacientes com diabetes mellitus na atenção primária à saúde: revisão integrativa. Braz J Hea Rev 2024; 7: e68794 [DOI: 10.34119/bjhrv7n2-331]
- Pantanetti P, Cangelosi G, Alberti S, Di Marco S, Michetti G, Cerasoli G, Di Giacinti M, Coacci S, Francucci N, Petrelli F, Ambrosio G, 6 Grinta R. Changes in body weight and composition, metabolic parameters, and quality of life in patients with type 2 diabetes treated with subcutaneous semaglutide in real-world clinical practice. Front Endocrinol (Lausanne) 2024; 15: 1394506 [PMID: 39015186 DOI: 10.3389/fendo.2024.1394506
- Chen J, Zhou H, Liu K. Normal weight and central obesity as predictors of increased all-cause mortality in metabolic dysfunction associated 7 steatotic liver disease. J Hepatol 2024; 80: e143-e145 [PMID: 38101754 DOI: 10.1016/j.jhep.2023.11.024]
- Park B, Yoon J, Tran TXM. Accounting for time-varying exposures and covariates in the relationship between obesity and diabetes: analysis 8 using parametric g-formula. J Epidemiol Community Health 2024 [PMID: 39025645 DOI: 10.1136/jech-2023-221882]
- Zhang H, Zhou XD, Shapiro MD, Lip GYH, Tilg H, Valenti L, Somers VK, Byrne CD, Targher G, Yang W, Viveiros O, Opio CK, Mantzoros 9 CS, Ryan JD, Kok KYY, Jumaev NA, Perera N, Robertson AG, Abu-Abeid A, Misra A, Wong YJ, Ruiz-Úcar E, Ospanov O, Kızılkaya MC, Luo F, Méndez-Sánchez N, Zuluaga M, Lonardo A, Al Momani H, Toro-Huamanchumo CJ, Adams L, Al-Busafi SA, Sharara AI, Chan WK, Abbas SI, Sookoian S, Treeprasertsuk S, Ocama P, Alswat K, Kong AP, Ataya K, Lim-Loo MC, Oviedo RJ, Szepietowski O, Fouad Y, Zhang H, Abdelbaki TN, Katsouras CS, Prasad A, Thaher O, Ali A, Molina GA, Sung KC, Chen QF, Lesmana CRA, Zheng MH. Global burden of metabolic diseases, 1990-2021. Metabolism 2024; 160: 155999 [PMID: 39151887 DOI: 10.1016/j.metabol.2024.155999]
- Ghusn W, Salame M, Sayegh L, Hage K, Storm AC, Dayyeh BKA, Ghanem OM. The association between microvascular and macrovascular 10 diseases and diabetes remission after bariatric surgery. Surg Endosc 2024; 38: 1835-1843 [PMID: 38305895 DOI: 10.1007/s00464-024-10687-0]
- Li S, Zhang P, Di J, Han X, Tu Y, Yang D, Xu R, Xiao Y, Zhou J, Bao Y, Yin J, Yu H, Jia W, Han J. Associations of change in body fat 11 percentage with baseline body composition and diabetes remission after bariatric surgery. Obesity (Silver Spring) 2024; 32: 871-887 [PMID: 38515375 DOI: 10.1002/oby.24003]
- 12 Courcoulas AP, Patti ME, Hu B, Arterburn DE, Simonson DC, Gourash WF, Jakicic JM, Vernon AH, Beck GJ, Schauer PR, Kashyap SR, Aminian A, Cummings DE, Kirwan JP. Long-Term Outcomes of Medical Management vs Bariatric Surgery in Type 2 Diabetes. JAMA 2024; 331: 654-664 [PMID: 38411644 DOI: 10.1001/jama.2024.0318]
- Yazici H, Ozturk AM, Cekic A, Yasar AC, Yildirim M. Laparoscopic Sleeve Gastrectomy as One-Step Procedure for Patients with Obesity: 13 Long-Term Outcomes of a Single Center. Bariatr Surg Pract P 2024; 19: 62-68 [DOI: 10.1089/bari.2023.0001]
- Gregg EW, Chen H, Bancks MP, Manalac R, Maruthur N, Munshi M, Wing R; Look AHEAD Research Group. Impact of remission from type 14 2 diabetes on long-term health outcomes: findings from the Look AHEAD study. Diabetologia 2024; 67: 459-469 [PMID: 38233592 DOI: 10.1007/s00125-023-06048-6]
- 15 Saberdoust F, Salehabadi G, Sheykholeslamy S, Noroozi E, Moradi M, Pazouki A, Kabir A. Diagnostic Value of Advanced-DiaRem for Predicting Diabetic remission after One Anastomosis Gastric Bypass/Minigastric Bypass. Obes Surg 2024; 34: 3467-3474 [PMID: 39080227 DOI: 10.1007/s11695-024-07431-x]
- Scott AW, Amateau SK, Leslie DB, Ikramuddin S, Wise ES. Rates and Risk Factors for 30-Day Morbidity After One-Stage Vertical Banded 16 Gastroplasty Conversions: A Retrospective Analysis. Am Surg 2024; 31348241248817 [PMID: 38641431 DOI: 10.1177/00031348241248817]
- Carbonaro J, McLaughlin T, Seip R, Staff I, Wu Y, Santana C, Bond D, Tishler D, Benbrahim A, Papasavas P. Five-year outcomes of 17 revisional bariatric surgery: gastric band to sleeve gastrectomy or to Roux-en-Y gastric bypass. Surg Endosc 2024; 38: 2719-2725 [PMID: 38532050 DOI: 10.1007/s00464-024-10764-4]
- 18 Papadia FS, Adami G, Razzetta A, Florenzano A, Longo G, Rubartelli A, Carlini F, De Cian O, Camerini G. Biliopancreatic diversion for severe obesity: long-term weight maintenance and occurrence of nutritional complications are two facets of the same coin. Br J Surg 2024; 111 [PMID: 38551118 DOI: 10.1093/bjs/znae058]
- Hedberg S, Thorell A, Österberg J, Peltonen M, Andersson E, Näslund E, Hertel JK, Svanevik M, Stenberg E, Neovius M, Näslund I, Wirén 19



WJD | https://www.wjgnet.com

M, Ottosson J, Olbers T; BEST Study Group. Comparison of Sleeve Gastrectomy vs Roux-en-Y Gastric Bypass: A Randomized Clinical Trial. JAMA Netw Open 2024; 7: e2353141 [PMID: 38289603 DOI: 10.1001/jamanetworkopen.2023.53141]

- 20 Winckelmann LA, Gribsholt SB, Bødkergaard K, Rejnmark L, Madsen LR, Richelsen B. Risk of fractures following bariatric surgery with Roux-en-Y gastric bypass or sleeve gastrectomy: a Danish population-based cohort study. Eur J Endocrinol 2024; 191: 1-8 [PMID: 38916995 DOI: 10.1093/ejendo/lvae068]
- Rodbard HW, Barnard-Kelly K, Pfeiffer AFH, Mauersberger C, Schnell O, Giorgino F. Practical strategies to manage obesity in type 2 21 diabetes. Diabetes Obes Metab 2024; 26: 2029-2045 [PMID: 38514387 DOI: 10.1111/dom.15556]
- Craig Wood G, Horwitz D, Still CD, Mirshahi T, Benotti P, Parikh M, Hirsch AG. Performance of the DiaRem Score for Predicting Diabetes 22 Remission in Two Health Systems Following Bariatric Surgery Procedures in Hispanic and non-Hispanic White Patients. Obes Surg 2018; 28: 61-68 [PMID: 28717860 DOI: 10.1007/s11695-017-2799-y]
- Hainer V, Toplak H, Mitrakou A. Treatment modalities of obesity: what fits whom? Diabetes Care 2008; 31 Suppl 2: S269-S277 [PMID: 23 18227496 DOI: 10.2337/dc08-s265]
- 24 Sanches E, Timmermans M, Topal B, Celik A, Sundbom M, Ribeiro R, Parmar C, Ugale S, Proczko M, Stepaniak PS, Pujol Rafols J, Mahawar K, Buise MP, Neimark A, Severin R, Pouwels S. Cardiac remodeling in obesity and after bariatric and metabolic surgery; is there a role for gastro-intestinal hormones? Expert Rev Cardiovasc Ther 2019; 17: 771-790 [PMID: 31746657 DOI: 10.1080/14779072.2019.1690991]
- Osto E, Doytcheva P, Corteville C, Bueter M, Dörig C, Stivala S, Buhmann H, Colin S, Rohrer L, Hasballa R, Tailleux A, Wolfrum C, Tona F, 25 Manz J, Vetter D, Spliethoff K, Vanhoutte PM, Landmesser U, Pattou F, Staels B, Matter CM, Lutz TA, Lüscher TF. Rapid and body weightindependent improvement of endothelial and high-density lipoprotein function after Roux-en-Y gastric bypass: role of glucagon-like peptide-1. Circulation 2015; 131: 871-881 [PMID: 25673670 DOI: 10.1161/CIRCULATIONAHA.114.011791]
- Faria G, Preto J, da Costa EL, Guimarães JT, Calhau C, Taveira-Gomes A. Acute improvement in insulin resistance after laparoscopic Roux-26 en-Y gastric bypass: is 3 days enough to correct insulin metabolism? Obes Surg 2013; 23: 103-110 [PMID: 23114971 DOI: 10.1007/s11695-012-0803-01
- Farb MG, Ganley-Leal L, Mott M, Liang Y, Ercan B, Widlansky ME, Bigornia SJ, Fiscale AJ, Apovian CM, Carmine B, Hess DT, Vita JA, 27 Gokce N. Arteriolar function in visceral adipose tissue is impaired in human obesity. Arterioscler Thromb Vasc Biol 2012; 32: 467-473 [PMID: 22095978 DOI: 10.1161/ATVBAHA.111.235846]
- Green DJ, Hopman MT, Padilla J, Laughlin MH, Thijssen DH. Vascular Adaptation to Exercise in Humans: Role of Hemodynamic Stimuli. 28 Physiol Rev 2017; 97: 495-528 [PMID: 28151424 DOI: 10.1152/physrev.00014.2016]
- Green DJ, Jones H, Thijssen D, Cable NT, Atkinson G. Flow-mediated dilation and cardiovascular event prediction: does nitric oxide matter? 29 Hypertension 2011; 57: 363-369 [PMID: 21263128 DOI: 10.1161/HYPERTENSIONAHA.110.167015]
- Tang HH, Wang D, Tang CC. Effect of bariatric surgery on metabolism in diabetes and obesity comorbidity: Insight from recent research. 30 World J Diabetes 2024; 15: 586-590 [PMID: 38680688 DOI: 10.4239/wjd.v15.i4.586]



WJD https://www.wjgnet.com



## Published by Baishideng Publishing Group Inc 7041 Koll Center Parkway, Suite 160, Pleasanton, CA 94566, USA Telephone: +1-925-3991568 E-mail: office@baishideng.com Help Desk: https://www.f6publishing.com/helpdesk https://www.wjgnet.com

