



Obesity Management Medications & Metabolic Surgery and the IFSO Consensus

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OSWALDO CRUZ
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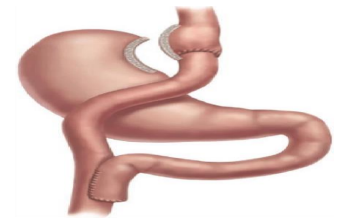
Disclosures

- Research Grant, Johnson & Johnson Medtech
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- SAB: Morphic Medical, Medtronic, Johnson & Johnson, Regeneron

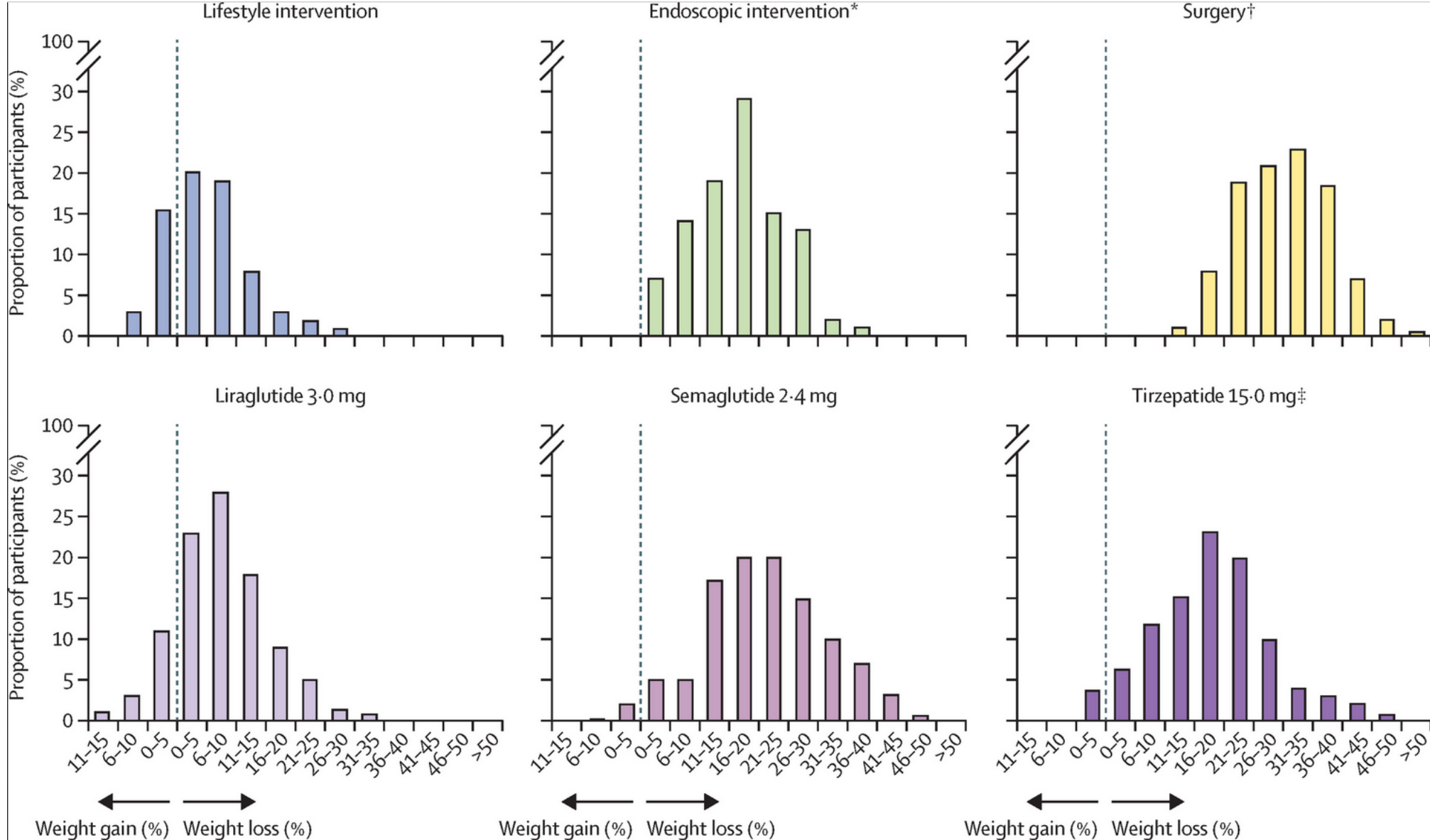


Bariatric/metabolic Surgery



- ✓ Long-term significant WL (Sjostrom L, JAMA, 2012)
- ✓ **Physiologic mechanisms** (Cummings D, D Care 2006)
- ✓ **Durable T2D remission** (glucocentric endpoint) (Courcoulas A, JAMA 2024)
- ✓ **Renoprotective** (Cohen RV, Lancet EC, 2022)
- ✓ **Decrease CV risk factors** (Sjostrom L, JAMA, 2012)
- ✓ **Decrease CV events and mortality** (Aminian A, JAMA, 2019)
- ✓ **Safe** (Arterburn D, JAMA 2022)

Heterogeneous response to obesity treatments



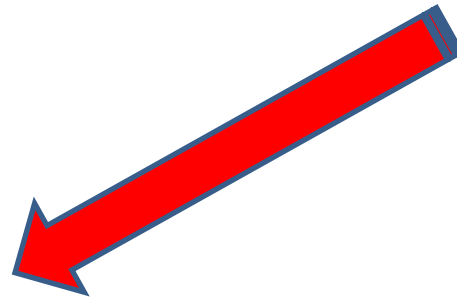
There is no magic bullet for obesity



*Ildiko Lingvay, Priya Sumithran,
Carel W le Roux, *Ricardo V Cohen*
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Lancet DE, July 2023

**DECISIONS ARE MORE IMPORTANT
THAN INCISIONS**



BEST OUTCOMES




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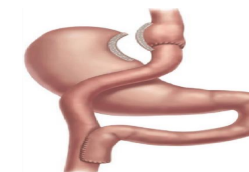
Metabolic surgery **&** **Pharmacotherapy**



Bariatric Surgery: There Is a Room for Improvement to Reduce Mortality in Patients with Type 2 Diabetes

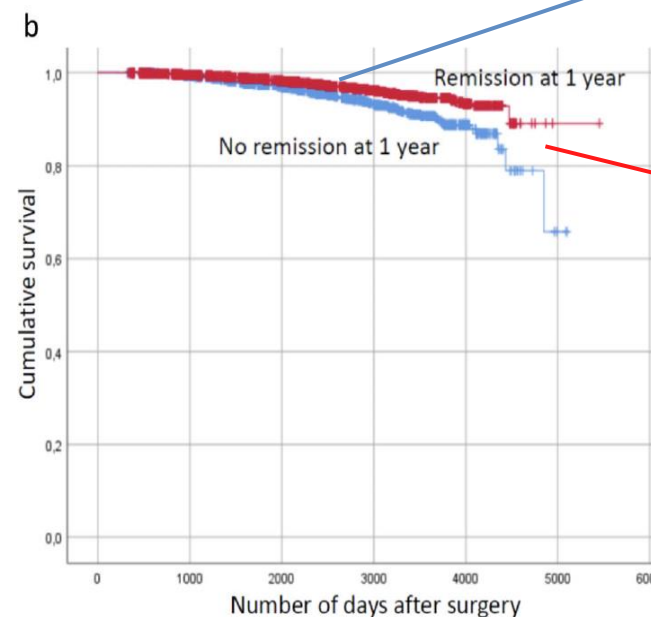
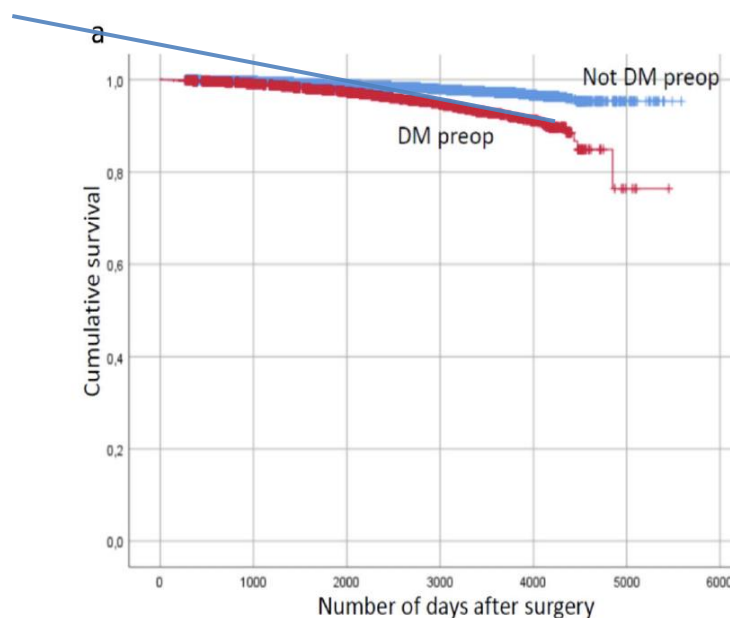
Carel W. le Roux¹ · Johan Ottosson^{2,3} · Erik Näslund^{2,4} · Ricardo V. Cohen⁵  · Erik Stenberg^{2,3} · Magnus Sundbom^{2,6} · Ingmar Näslund^{2,3}

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RYGB pts with T2D have higher mortality during FU than without T2D

SoReg, Scandinavian Obesity Surgery Registry 65,345 pts up to 10y FU, all after RYGB



Adjunctive pharmacotherapy

Lower mortality if T2D remission @1y

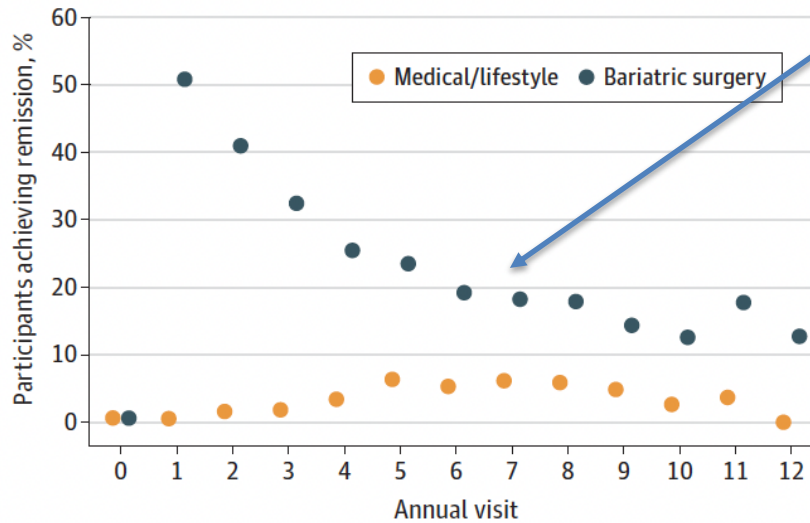
Long-Term Outcomes of Medical Management vs Bariatric Surgery in Type 2 Diabetes

Anita P. Courcoulas, MD; Mary Elizabeth Patti, MD; Bo Hu, PhD; David E. Arterburn, MD; Donald C. Simonson, MD, ScD; William F. Gourash, PhD; John M. Jakicic, PhD; Ashley H. Vernon, MD; Gerald J. Beck, PhD; Philip R. Schauer, MD; Sangeeta R. Kashyap, MD; Ali Aminian, MD; David E. Cummings, MD; John P. Kirwan, PhD

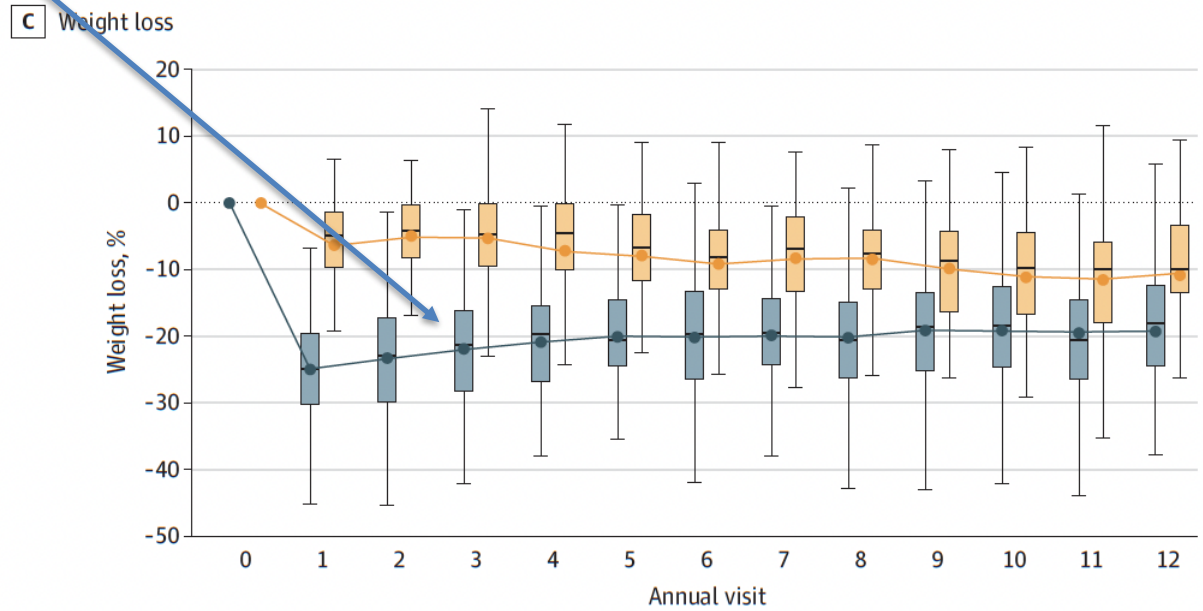
2024

Addition of pharmacotherapy

Figure 3. Diabetes Remission



No. of participants													
Medical/lifestyle	96	92	87	82	78	84	76	79	72	70	67	55	31
Bariatric surgery	166	164	151	149	140	146	108	131	116	125	117	99	82



No. at risk													
Bariatric surgery	166	164	161	158	144	149	122	139	121	126	121	106	85
Medical/lifestyle	96	91	84	86	79	78	77	75	73	73	70	60	34

Remission was defined as hemoglobin A_{1c} less than 6.5% and not receiving any medications for diabetes.






Collaborative Research

BJS, 2024, znae283

<https://doi.org/10.1093/bjs/znae283>

Collaborative Research Proceedings

International consensus position statement on the role of obesity management medications in the context of metabolic bariatric surgery: expert guideline by the International Federation for the Surgery of Obesity and Metabolic Disorders (IFSO)

Ricardo V. Cohen^{1,*} , Luca Busetto² , Randy Levinson³, Carel W. Le Roux⁴, Paulina Salminen^{5,6}  and Gerhard Prager⁷ on behalf of the experts of the International Consensus on the Role of Obesity Management Medications in the Context of Metabolic Bariatric Surgery

OMMs before surgery

Controversial data

OMMs **before** MBS

	Grade	Consensus (%)	Nr.of rounds	Nr.of total votes
There is insufficient high-level evidence to recommend the routine use of OMMs for weight loss before MBS	A+	100	2	37

Preoperative WL may influence perioperative outcomes

Preoperative weight loss is linked to improved mortality and leaks following elective bariatric surgery: an analysis of 548,597 patients from 2015–2018

Valentia Mocanu, M.D.¹, Gabriel Marci, M.D., Jerry T. Dang, M.D., Daniel W. Birch, M.D., M.Sc., Nishu J. Switzer, M.D., M.P.H., Shahzeer Karmali, M.D., M.P.H.

Department of Surgery, University of Alberta, Edmonton, Alberta, Canada
Received 2 March 2021; accepted 29 June 2021

Preoperative Weight Loss as a Predictor of Bariatric Surgery Postoperative Weight Loss and Complications

Jamil S. Samaan¹ · Jasmine Zhao² · Elaine Qian² · Angelica Hernandez² · Omar Toubat² · Evan T. Alicuben² · Yousaf Malik² · Kulmeet Sandhu² · Adrian Dobrowolsky² · Kamran Samakar²

Journal of Gastrointestinal Surgery 26.1 (2022): 86-93.

Preoperative WL does not influence perioperative outcomes and delays the operation

Preoperative weight loss: is waiting longer before bariatric surgery more effective?

Victor Eng, B.S.¹, Luis Garcia, M.S.¹, Habib Khoury, B.S.¹, John Morton, M.D., M.P.H.², Dan Azagury, M.D.^{1,2}

¹Bariatric and Minimally Invasive Surgery, Stanford School of Medicine, Stanford, California
²David Geffen School of Medicine, University of California at Los Angeles, Los Angeles, California
Received 18 April 2019; accepted 5 March 2019

Surgery for Obesity and Related Diseases 15 (2019) 951–957

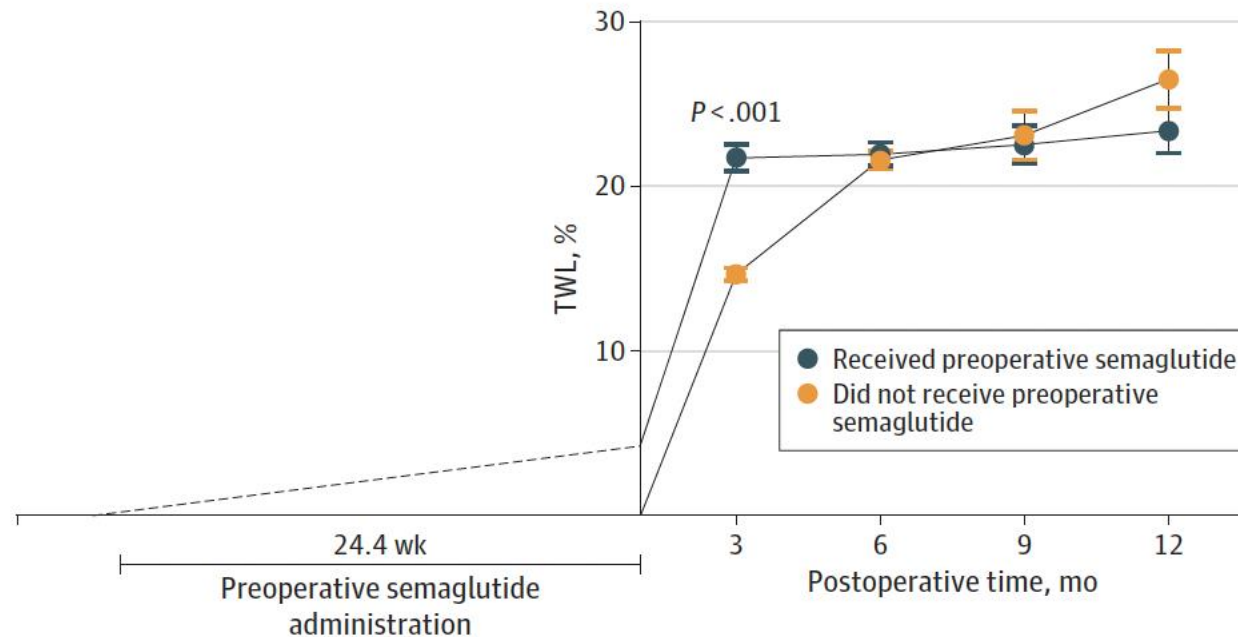
RESEARCH LETTER

Neoadjuvant Semaglutide, Bariatric Surgery Weight Loss, and Overall Outcomes

JAMA Surgery Published online March 5, 2025

Vasundhara Mathur, MD
Katherine Wasden, BA
Thomas H. Shin, MD, PhD
Pourya Medhati, MD
Abdelrahman A. Nimeri, M
Ali Tavakkoli, MD
Eric G. Sheu, MD, PhD

B Surgical and medical TWL



Neoadjuvant semaglutide:

- ✓ No WL benefits
- ✓ No greater intraop safety
- ✓ Surgical weight loss was significantly lower in patients treated with neoadjuvant semaglutide

OMMs **after** MBS

	Grade	Consensus (%)	Nr.of rounds	Nr.of total votes
Emerging evidence indicates that the weight loss induced by OMMs is similar among people who have or have not undergone MBS	A+	100	2	36

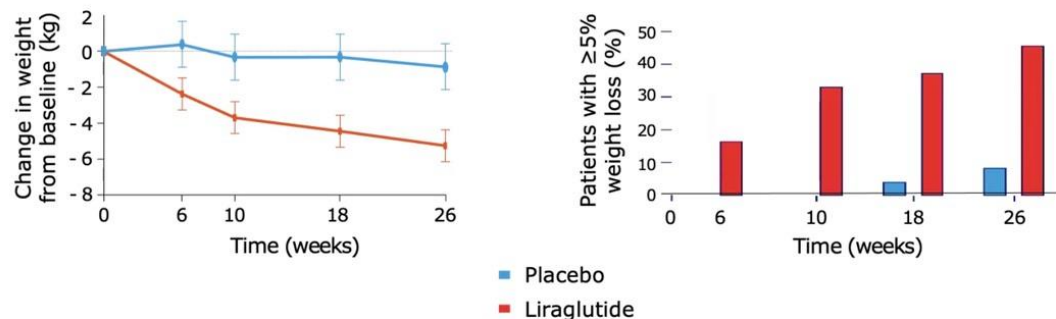
Mok et al, 2023

The BARI-OPTIMISE Randomized Clinical Trial

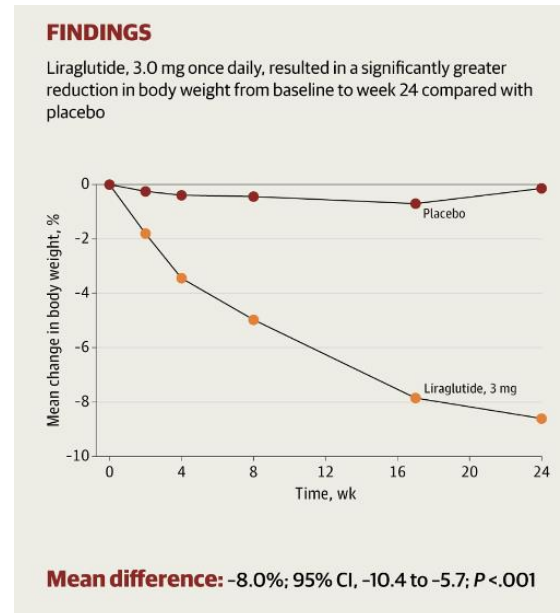
Postoperative pharmacotherapy augments surgical weight loss

GRAVITAS Study

Liraglutide after Gastric Bypass in T2D



Miras AD et al., Lancet Diabetes Endocrinol 2019

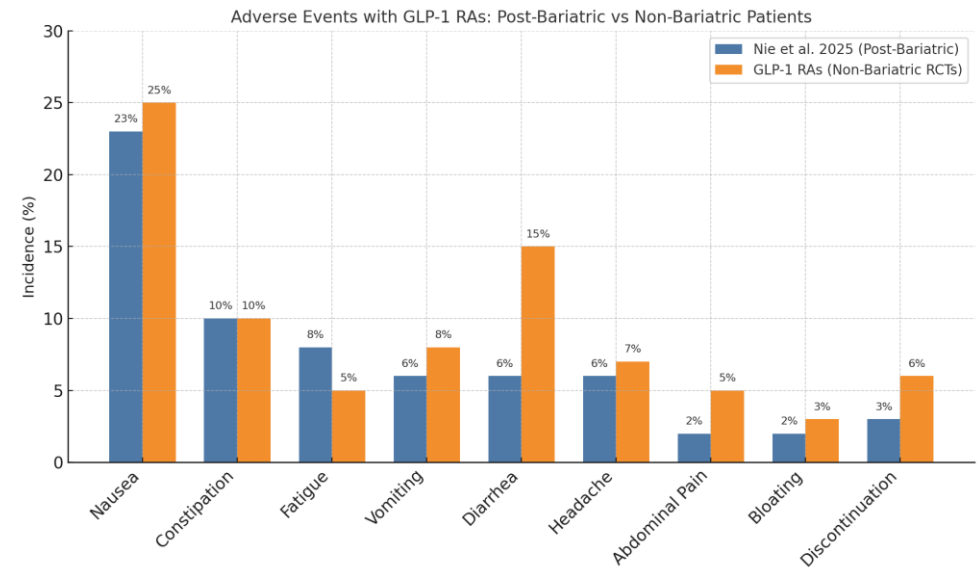
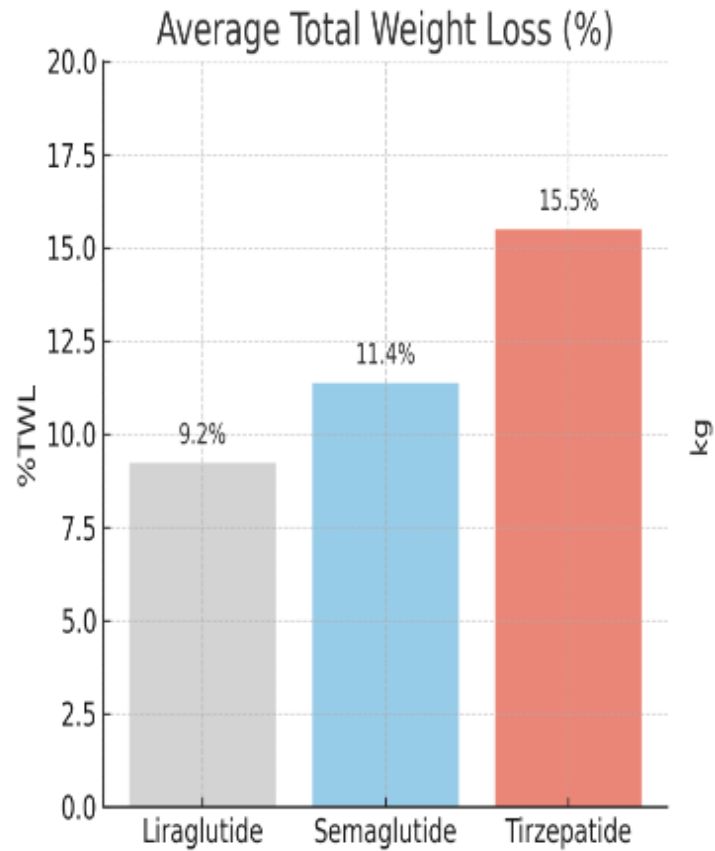


Cohen RV et al, BJS, 2024



Glucagon-Like Peptide-1 Receptor Agonists for the Treatment of Suboptimal Initial Clinical Response and Weight Gain Recurrence After Bariatric Surgery: a Systematic Review and Meta-analysis

Yuntao Nie¹ · Yiran Zhang² · Baoyin Liu¹ · Hua Meng¹



Indirect comparison with Step1 and Surmount 1 studies

Real world efficacy of naltrexone/bupropion for weight management in obesity and after bariatric surgery

Sept 2025

Marie Yskout¹, Nele Steenackers^{2,3}, Jarne Hoste⁴, Sofia Pazmino², Caroline Simoens², Nele Mattelaer², Ellen Deleus^{2,5}, Matthias Lannoo^{2,5}, Ann Mertens^{1,2}, Bart Van der Schueren^{1,2} and Roman Vangoitsenhoven^{1,2}✉

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International Journal of Obesity

111 surgery-naive patients (72.5%)

40 post-MBS patients (26.1%)


Study Overview

Predominantly female (77.5% surgery-naive, 75.0% post-MBS)

Median age: 45.0 years (surgery-naive), 51.5 years (post-MBS)

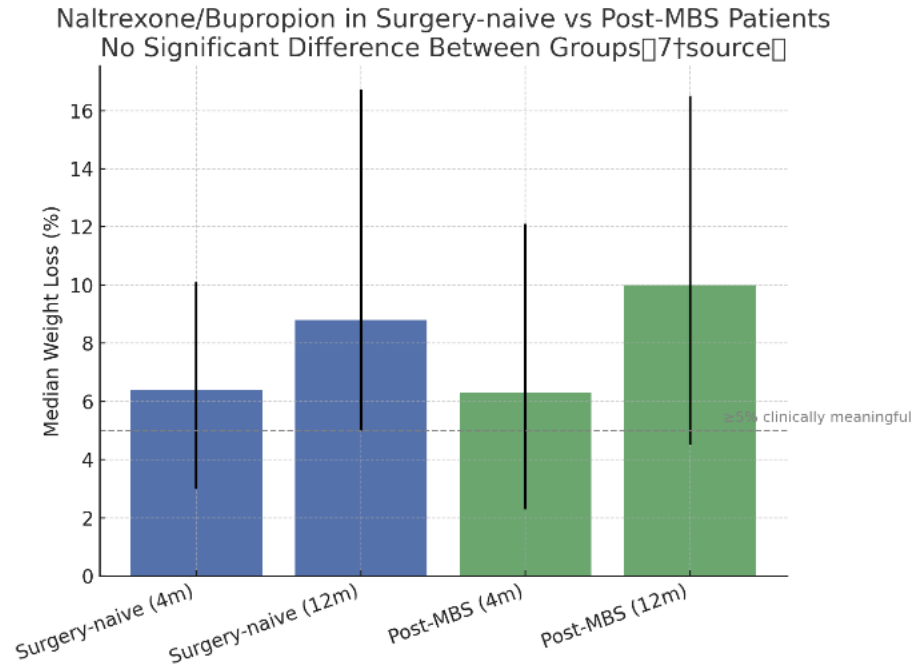
Treatment Protocol & Dosing

- **Starting dose:** 8mg naltrexone/90mg bupropion daily (one tablet)
- **Target dose:** 32mg naltrexone/360mg bupropion daily (two tablets twice daily)
- **Escalation:** Weekly increases per manufacturer protocol
- **Flexibility:** Submaximal doses allowed for:
 - Adverse effects management
 - Sufficient therapeutic response

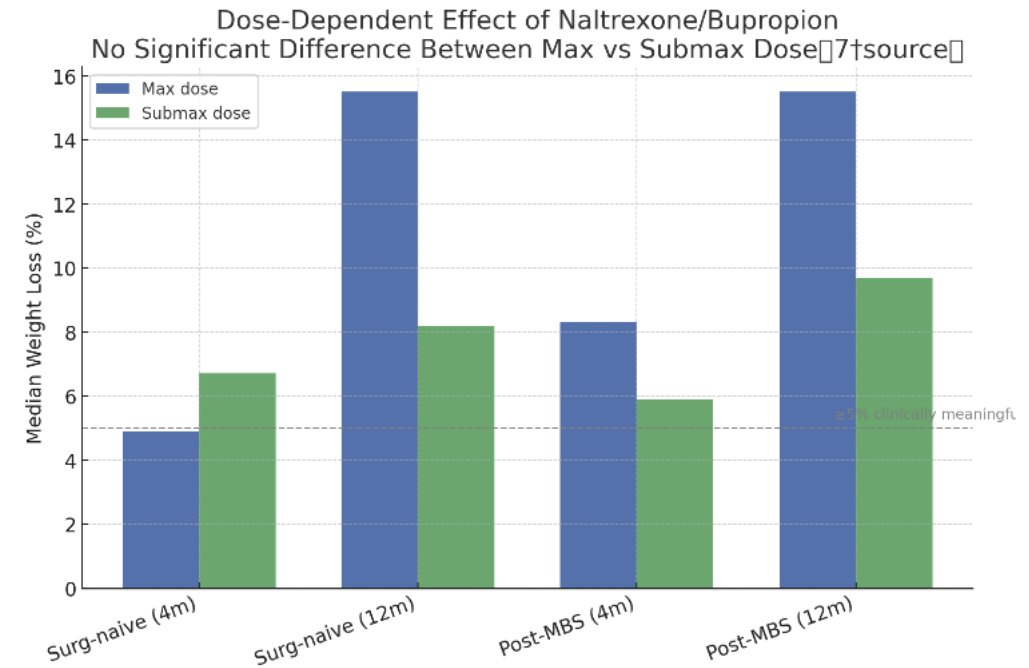
	Week 1	Week 2	Week 3	Week 4 and Beyond
 AM: Take with breakfast	 1 tablet	 1 tablet	 2 tablets	 2 tablets
 PM: Take before dinner		 1 tablet	 1 tablet	 2 tablets

Follow-up visits were scheduled at 4 and 12 months after treatment initiation

Naltrexone/Bupropion in Surgery-naive vs Post-MBS Patients...



Dose-Dependent Effect of Naltrexone/Bupropion...

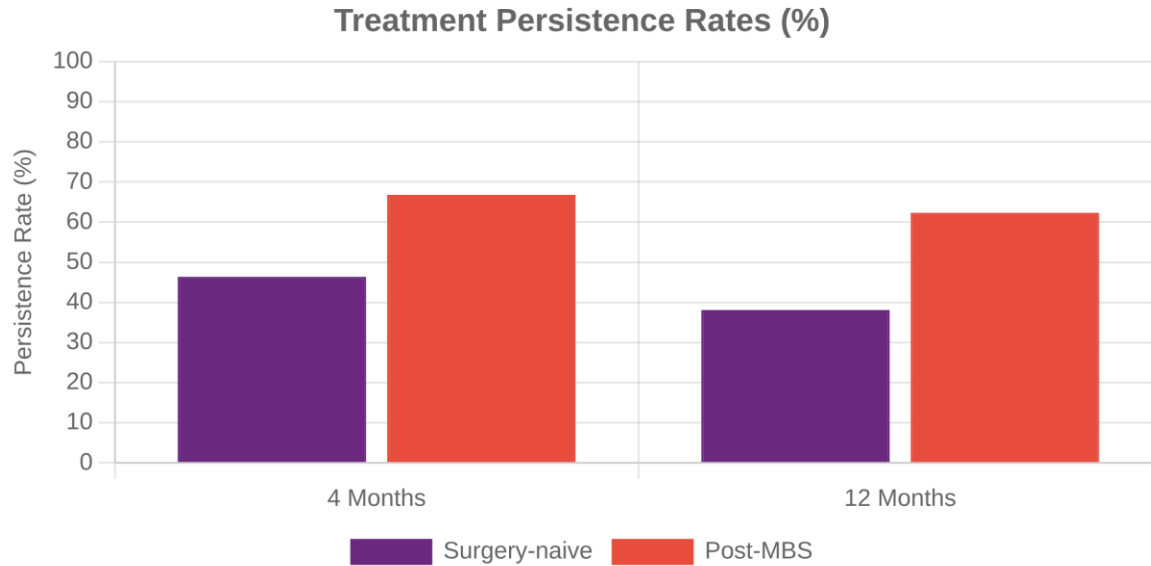
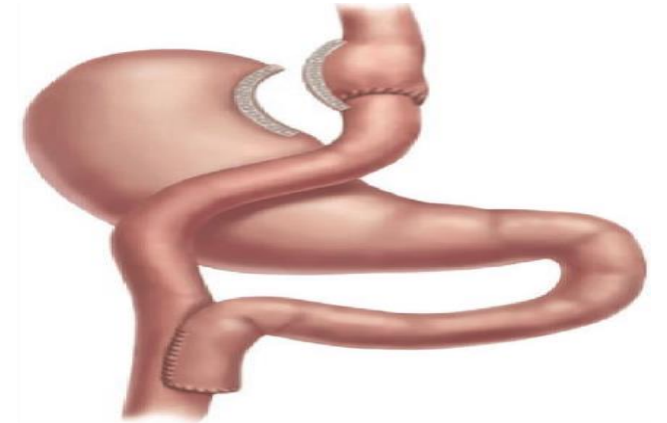


No dose-dependent effect

Sufficient Subjective Effect (Primary reason)

Surgery-naive: **62.1%** Post-MBS: **63.6%**

Treatment Persistence Rates



● 4-Month Persistence:

Surgery-naive: 46.9% (52/111 patients)

Post-MBS: 67.5% (27/40 patients)

● 12-Month Persistence:

Surgery-naive: 38.5% (20/52 patients)

Post-MBS: 63.0% (17/27 patients)

Key Finding:

Post-MBS patients showed significantly higher treatment persistence rates at both time points

OMMs **after** MBS

When to start

	Grade	Consensus (%)	Nr.of rounds	Nr.of total votes
Treatments with OMMs after MBS should generally be withheld until the achievement of weight plateau unless there is a compelling clinical need for earlier initiation	A+	100	3	33

Disclosing the effects of MBS before the introduction of OMMs

OMMs **after** MBS (OMMs / revisional surgery)

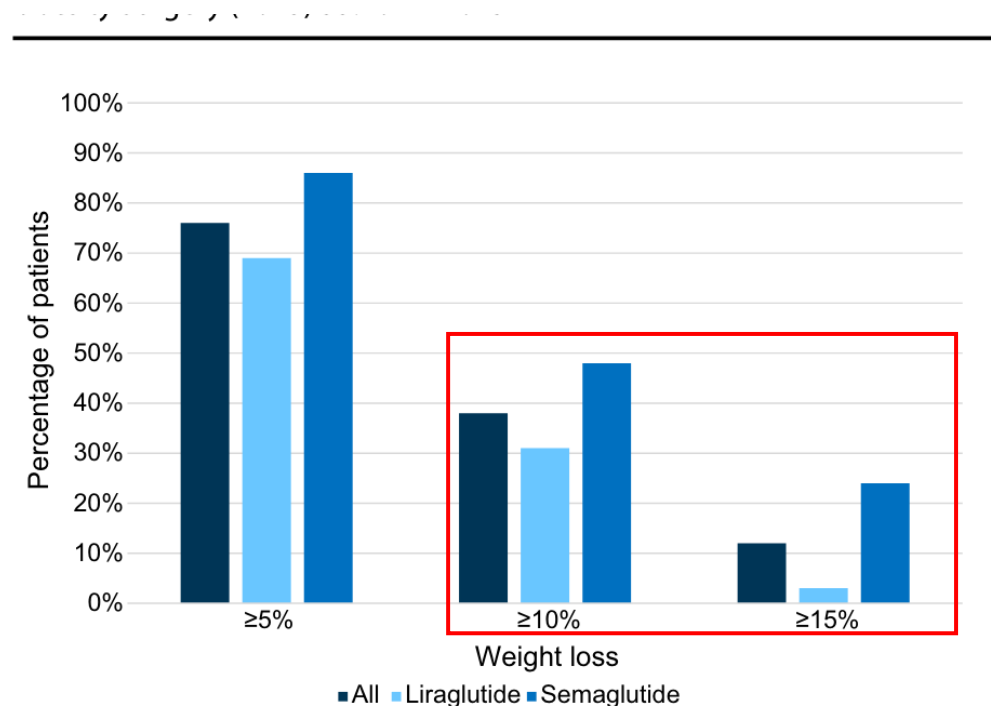
	Grade	Consensus (%)	Nr. of rounds	Nr. of total votes
For patients with recurrent weight gain, treatment with available OMMs should be considered prior to revisional surgery.	A	92	1	38

- ✓ Revisional MBS = higher morbidity, no robust data on outcomes
- ✓ Excellent safety profile and efficacy of OMMs

X

WL with GLP1RA for RWG/SoCR

Jensen, 2023

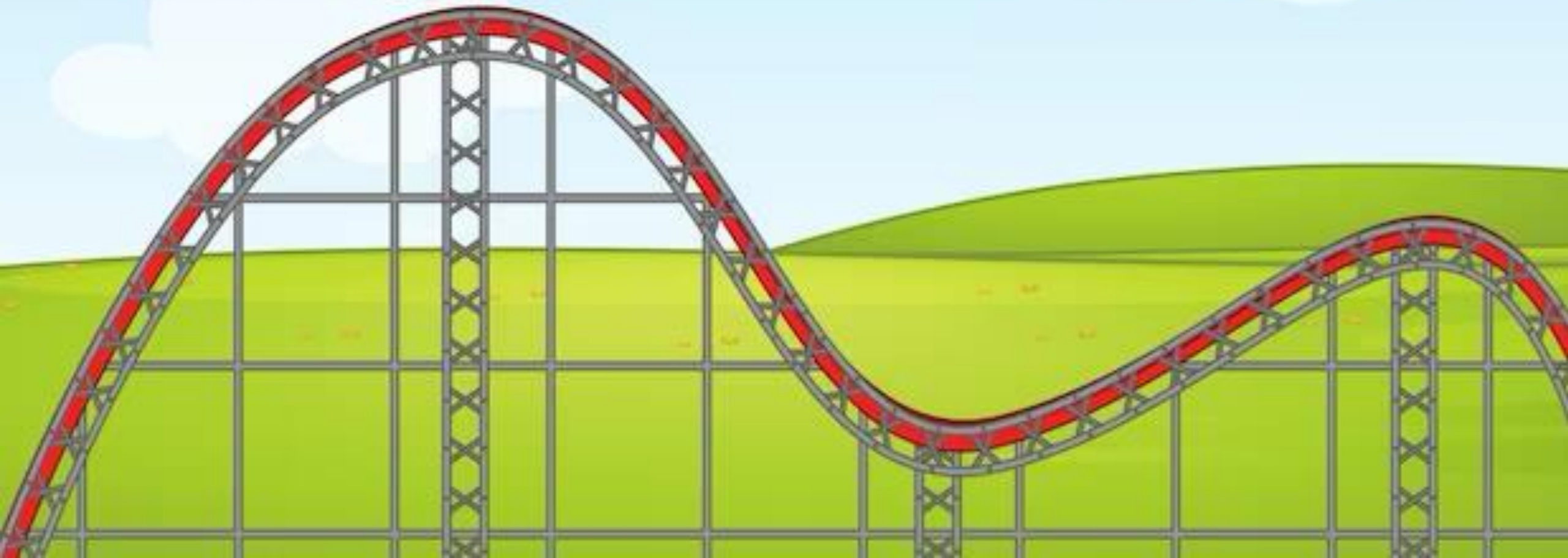


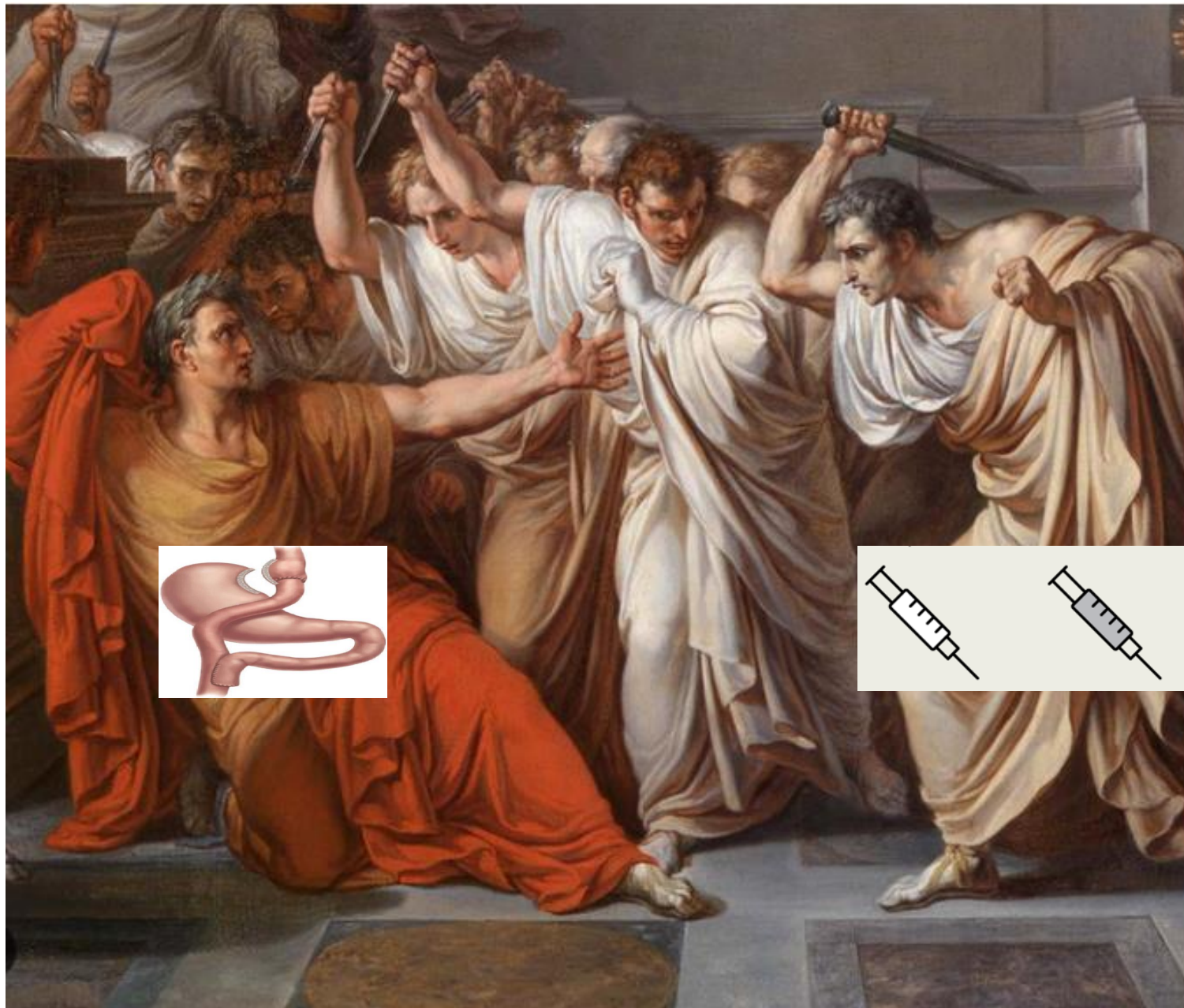
Pending questions of modern pharmacotherapy



- **Real world**
- Long-term adherence
 - \$\$
 - Access
 - Polypharmacy
 - Tolerability (short and long-term 27% de dropout in Select)
- Long-term efficacy
- Extreme obesity

How modern pharmacotherapy may affect surgical indications?





The Death of Julius Caesar (1806) by Vincenzo Camuccini in the National Museum of Capodimonte, in Naples

Metabolic surgery in era of modern pharmacotherapy

- ✓ Advancements in CV medications and PCI did not kill CABG
- ✓ Advancements in chemo, radio and immunotherapy did not kill cancer surgery
- ✓ Multifactorial disease, no magic bullet

A photograph of a long, straight road stretching towards a bright sunset on the horizon. The road is flanked by dark, silhouetted hills. The sky is filled with soft, white clouds. The sun is a bright, glowing orb just above the horizon, creating a lens flare effect. The road surface is dark asphalt with white lane markings. Large, white, stylized numbers representing years are painted on the road, starting from the foreground and receding into the distance. The years are 2023, 2024, 2025, 2026, and 2027, arranged in a sequence that follows the perspective of the road.

2023
2024
2025
2026
2027

Candidates in the new obesity treatment era

Pts preference

Extreme Obesity BMI > 45-50

Suboptimal response to med tx

Intolerance to pharmacotherapy

Candidates in
the new
obesity
treatment
era

Contraindications to
medicines

Cost

International Federation for the Surgery of Obesity statement on metabolic bariatric surgery after pharmacotherapy-induced weight loss in clinical obesity

Lancet Diabetes Endocrinol 20

Published Online

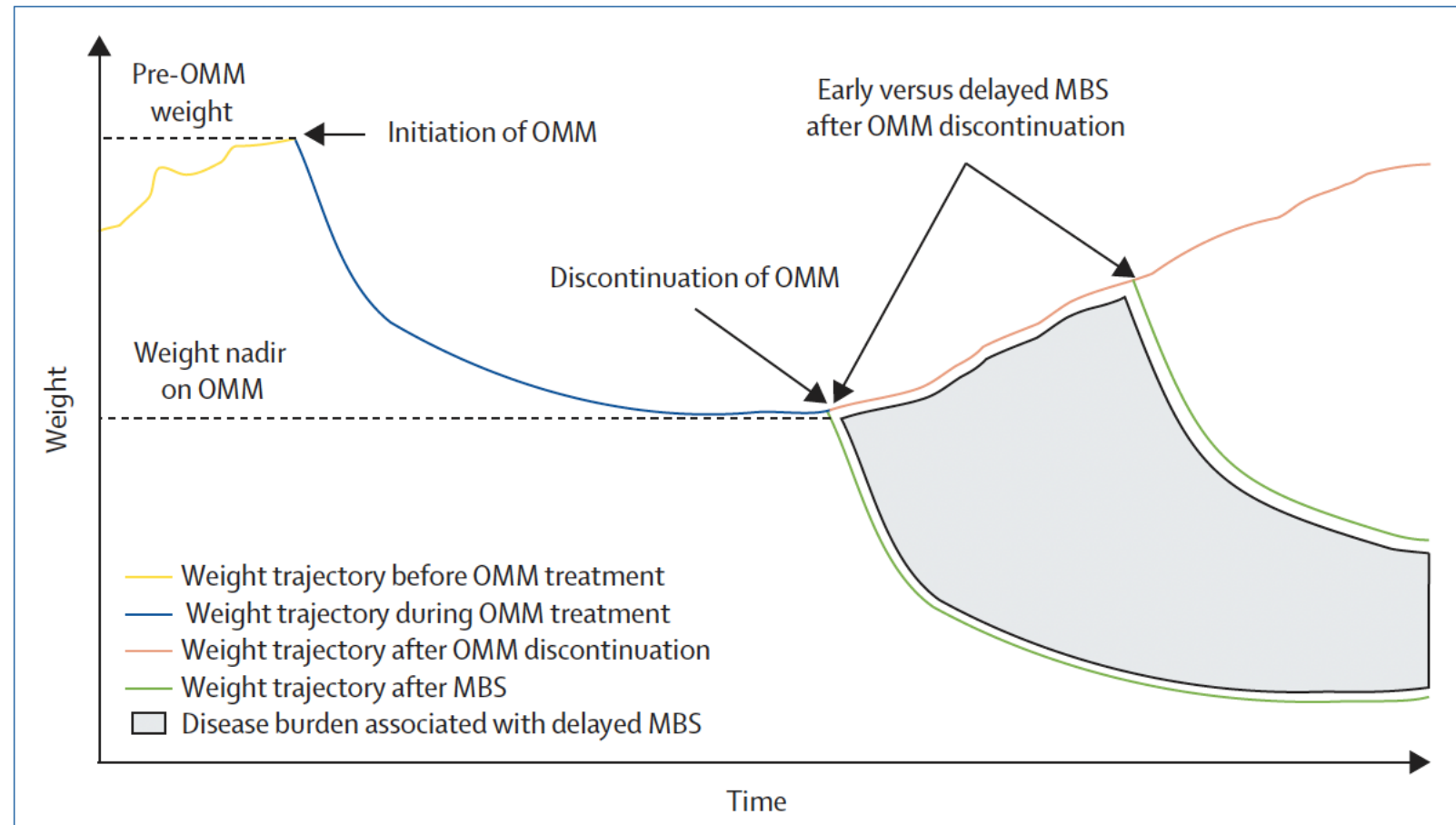
July 22, 2025

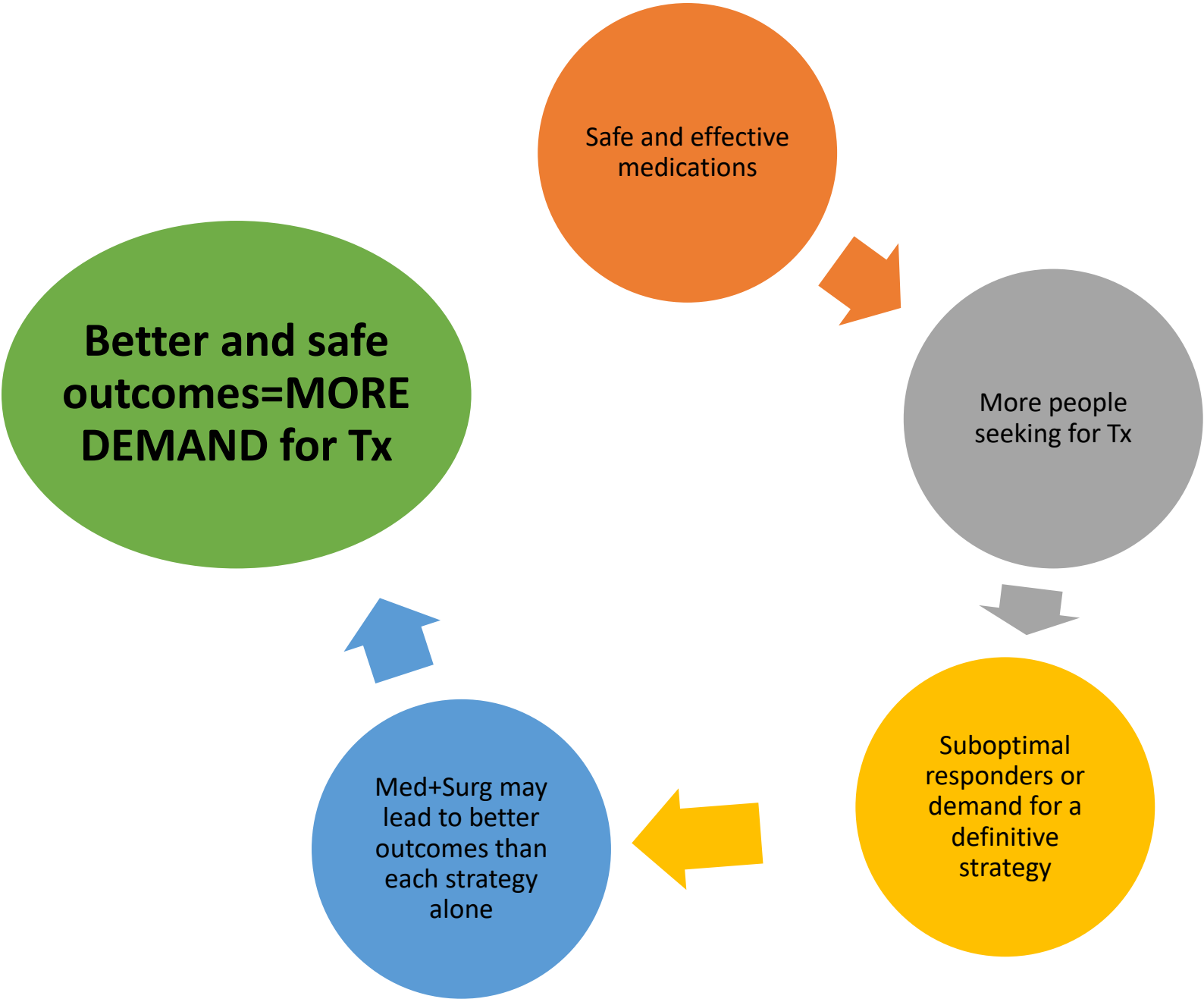
[https://doi.org/10.1016/](https://doi.org/10.1016/S2213-8587(25)00198-6)

[S2213-8587\(25\)00198-6](https://doi.org/10.1016/S2213-8587(25)00198-6)

*Ricardo V Cohen, Gerhard Prager, Carel W le Roux,
Ildiko Lingvay, Paulina Salminen
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- Good outcomes, however:
- Lack of access
- Long-term side effects
- Cost of continued Tx
- "I don't want it anymore"





MBS TODAY

Oncology model
What would an oncologist do ?



Access to full spectrum of therapy

Most invasive

Reoperative therapy

Adjuvant therapy

Surgical treatment

Combination therapy, when needed

Least invasive

Neoadjuvant therapy

An aerial photograph of a city, likely São Paulo, Brazil, showing a dense urban landscape with numerous high-rise buildings. In the foreground, a large, modern white building complex with a grid-like facade is prominent. The sky is clear and blue.

Thank you

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