

Maximizing Long-Term
Surgical Success:

The Emerging Role of Obesity Medications



Featured faculty




Richard Cohen, MD, PhD

President, IFSO
Director, The Center for the Treatment of Obesity
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Oswaldo Cruz German Hospital
Sao Paulo, Brazil



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Associate Professor of Medicine
Director, Nutrition and Obesity Research Program
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Rochester, Minnesota, USA

A person with long hair, wearing a red long-sleeved shirt, is seated in a black wheelchair. They are positioned in the center of a wide, open doorway, looking out towards a bright sunset. The sun is low on the horizon, creating a strong backlight effect and casting a warm glow. The sky is a mix of blue and orange. The interior of the building is visible on either side of the doorway, showing dark wood paneling and a light-colored floor. The overall mood is contemplative and hopeful.

Precision Obesity and Bariatrics: The Next Frontier

Andres Acosta, MD, PhD

Metabolic and Bariatric Surgery: Pathways to Better Outcomes

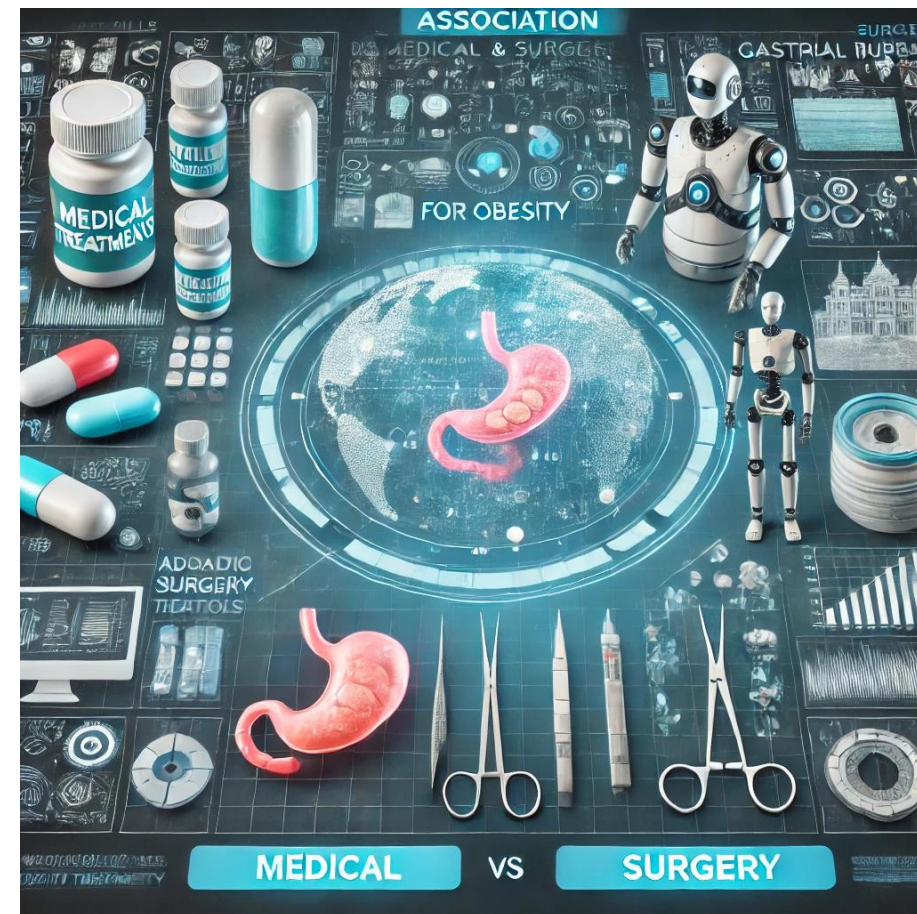
RICARDO COHEN

Director, The Center for the treatment of Obesity and Diabetes Hospital Alemao Oswaldo Cruz, São Paulo Brazil

President IFSO Global

Past President SBCBM (2011-2012)

Past President IFSO LAC (2018-2019)

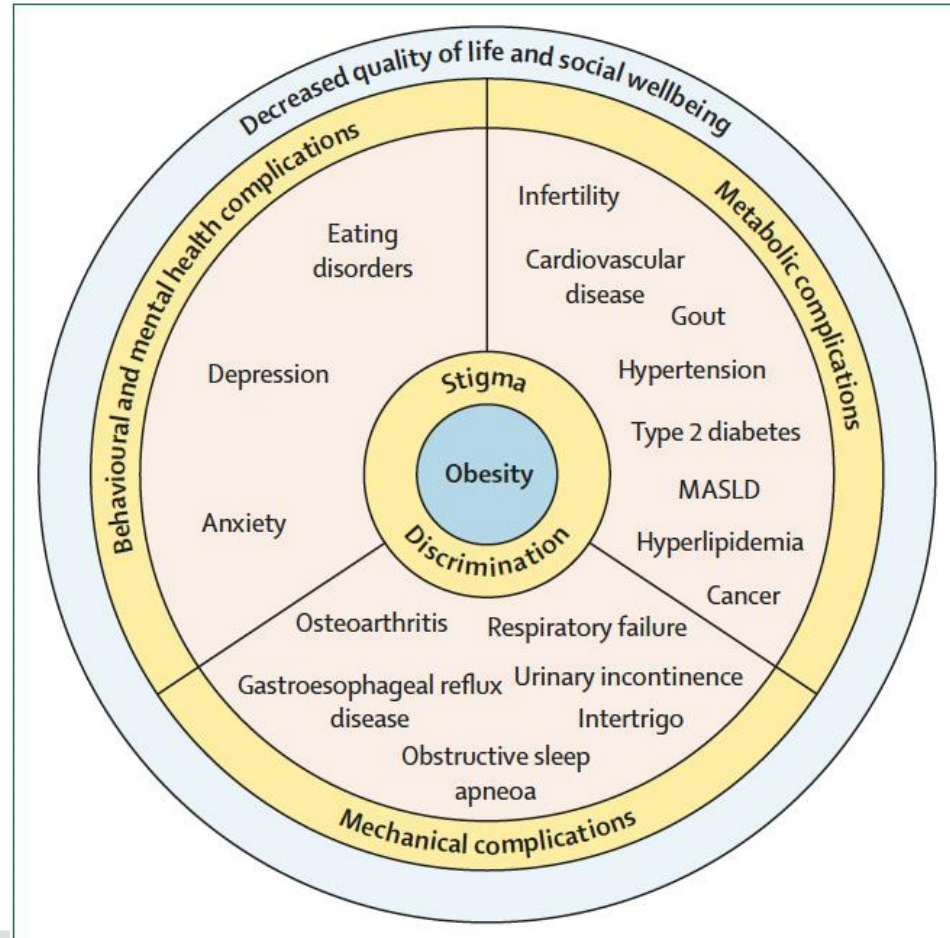


Disclosures

- Research Grant, Johnson & Johnson Medtech
- Research Grant, Medtronic
- Research Grant, GI Dynamics
- Research Grant, Hospital Oswaldo Cruz Bioscience Institute
- Research Grant, Marlex, Brazil
- Paid lectures, Johnson & Johnson Medtech, Medtronic, NovoNordisk, Merck, Boston Scientific

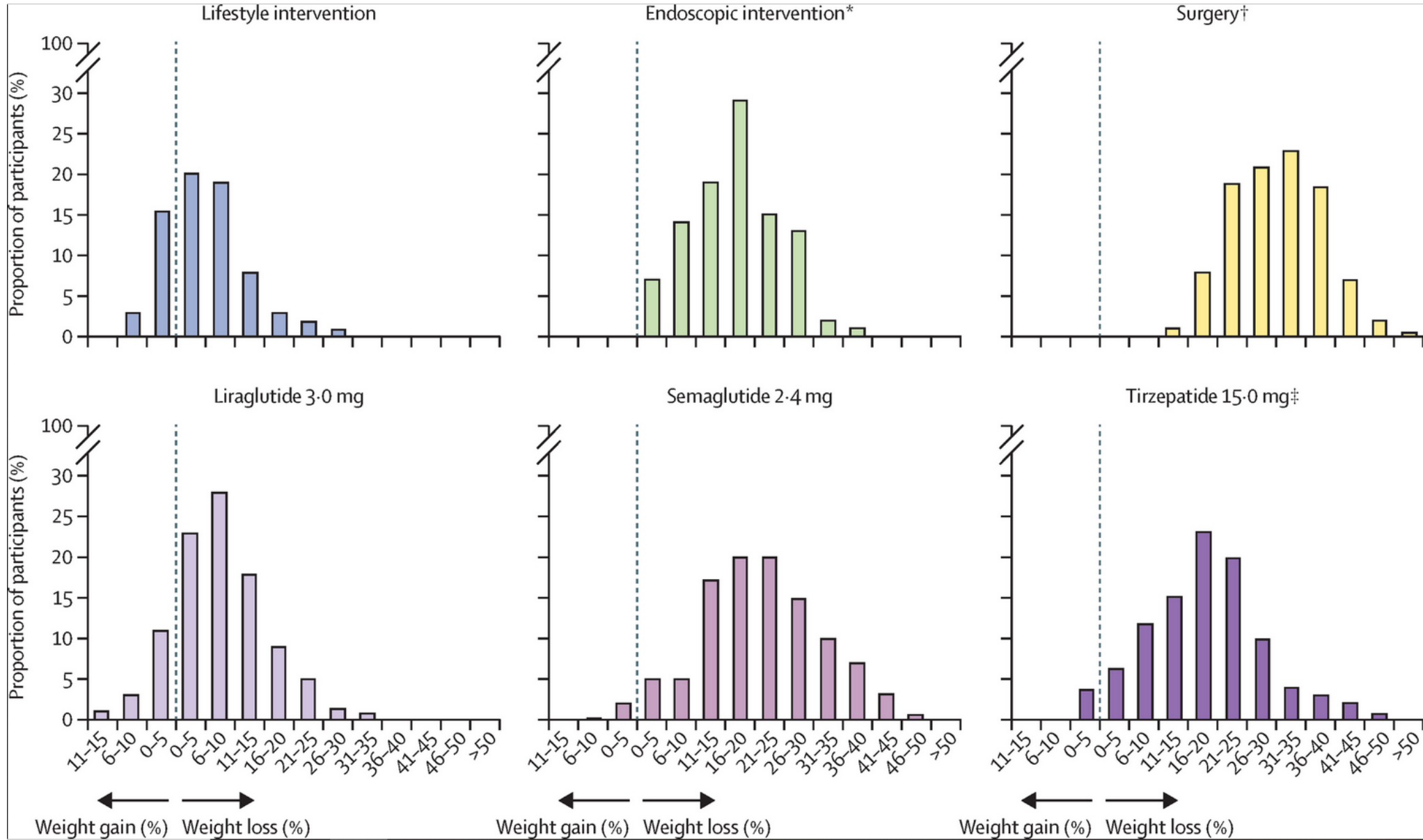
- SAB: Morphic Medical, Medtronic, Johnson & Johnson, Regeneron

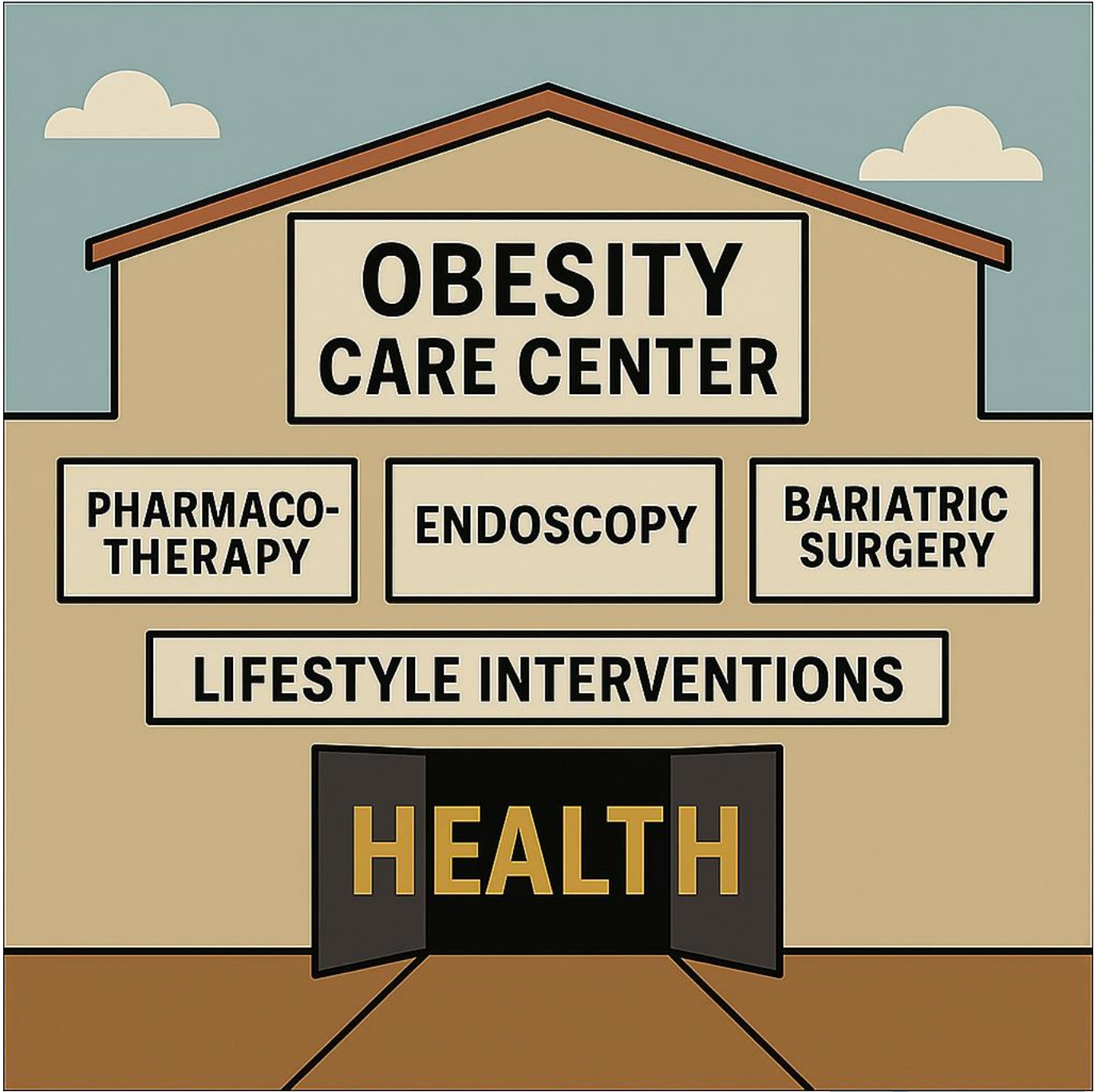
Obesity Is Not Just “Weight”



Clinical obesity is a multifaceted complex disease with signs, symptoms and complications

Heterogeneous response to obesity treatments



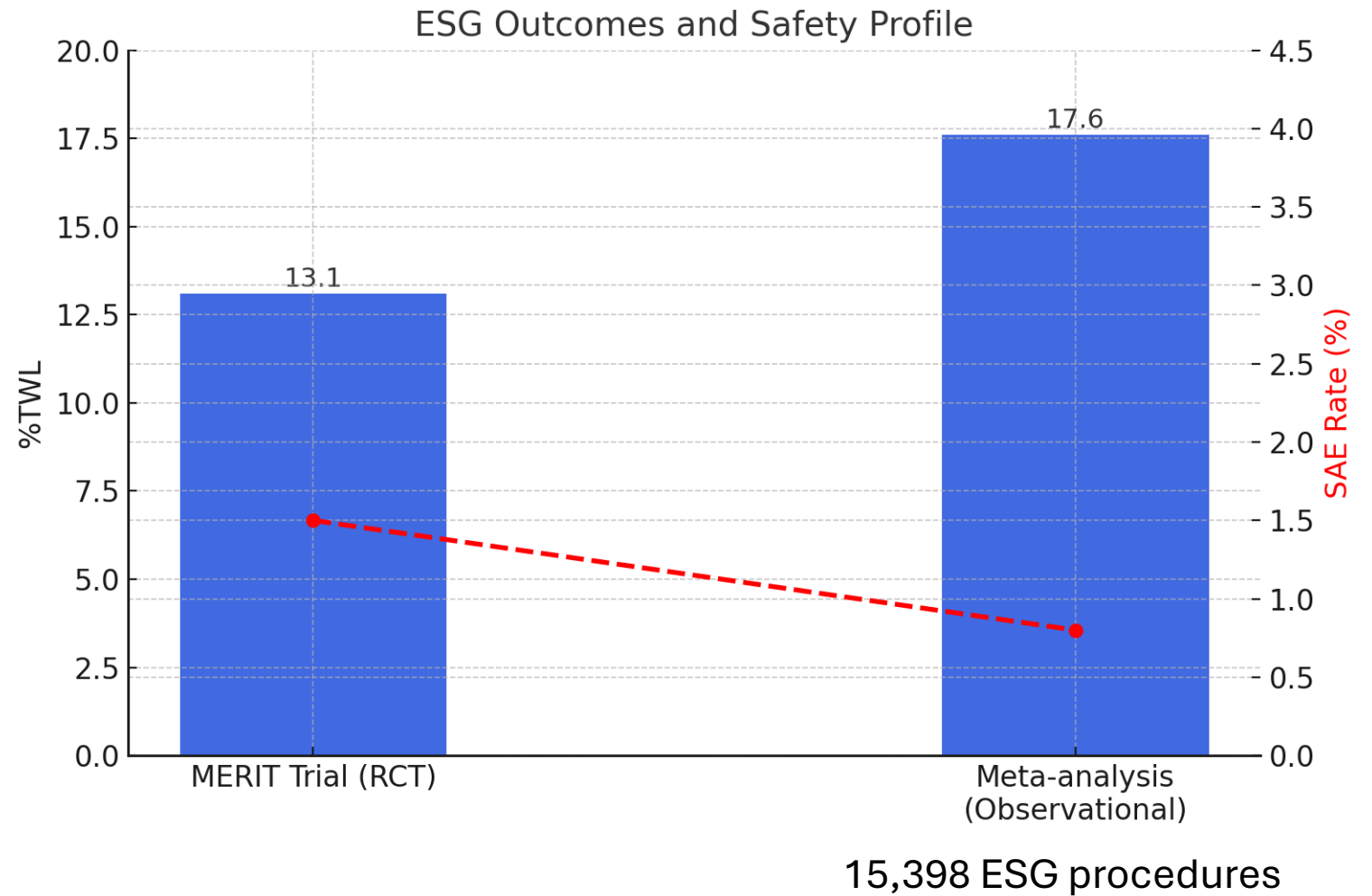
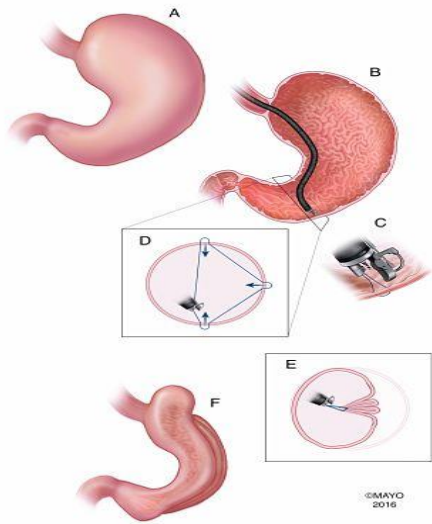


**PHARMACO-
THERAPY**

**ENDOSCOPIC
PROCEDURES**

**METABOLIC
BARIATRIC
SURGERY**

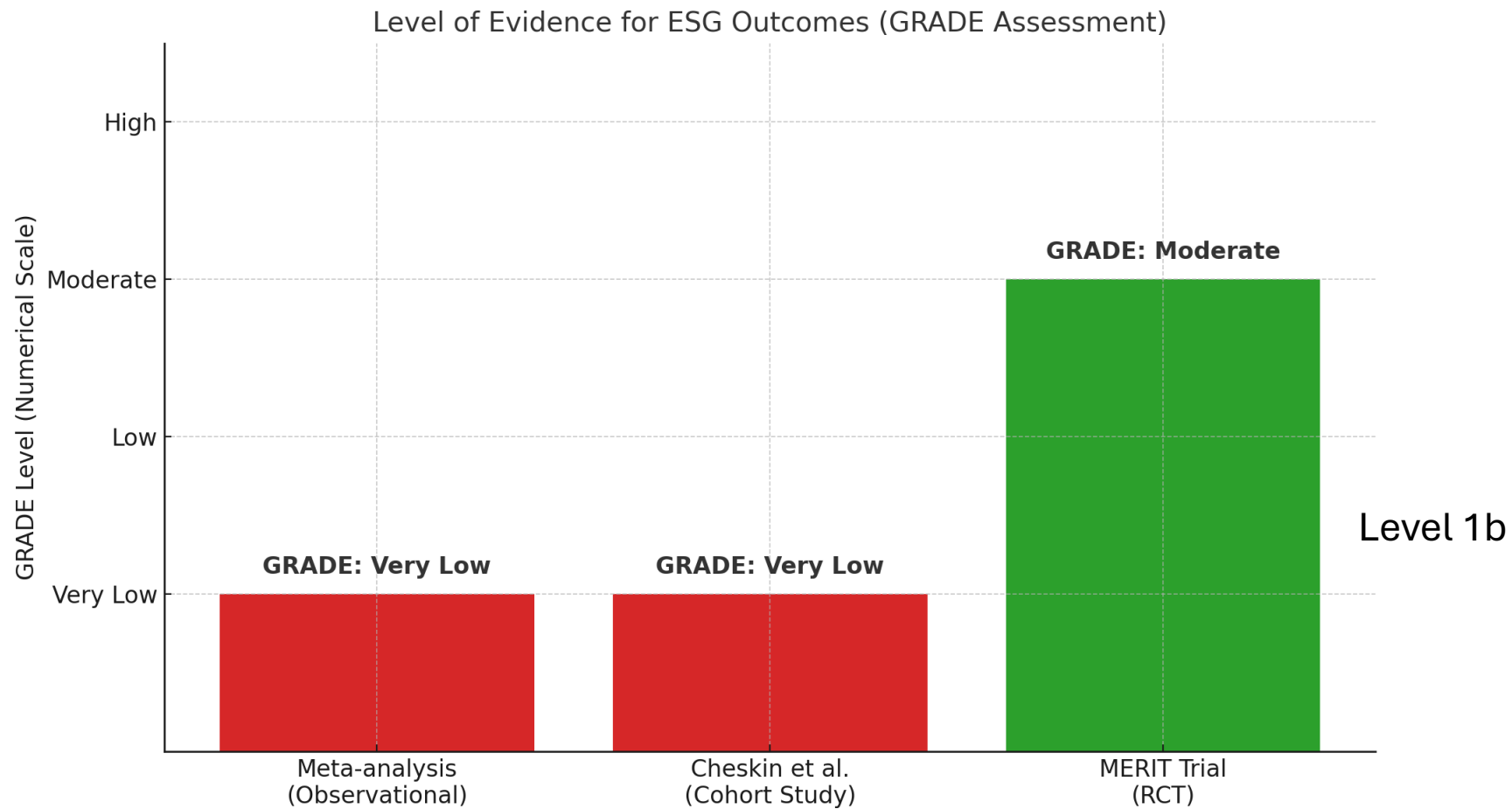
Endoscopic Sleeve Gastroplasty

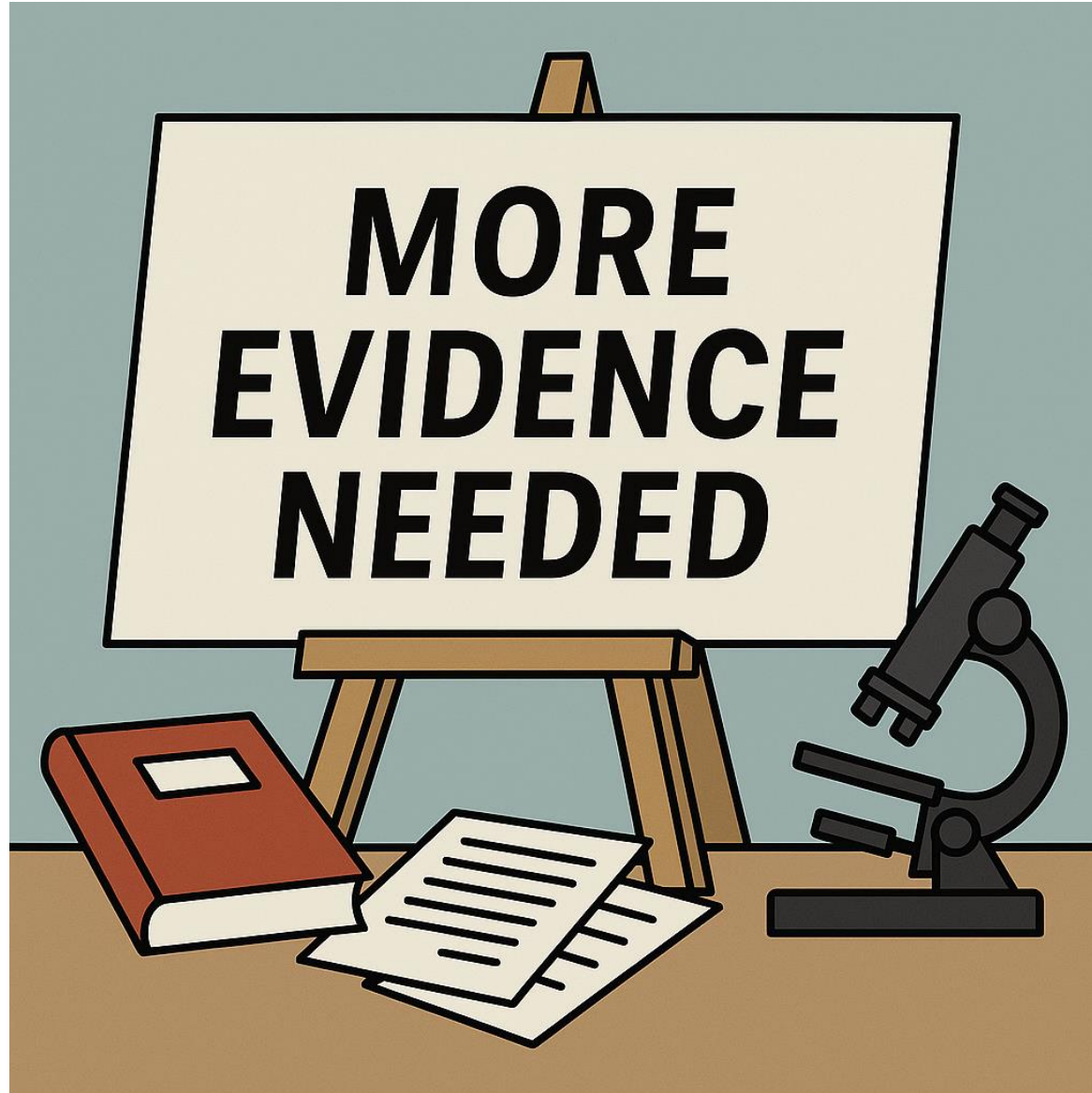
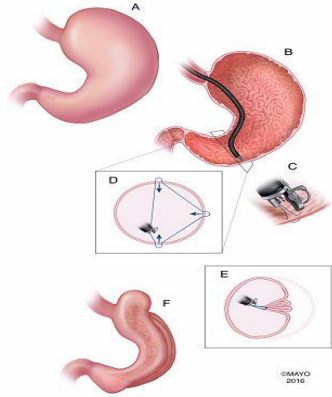


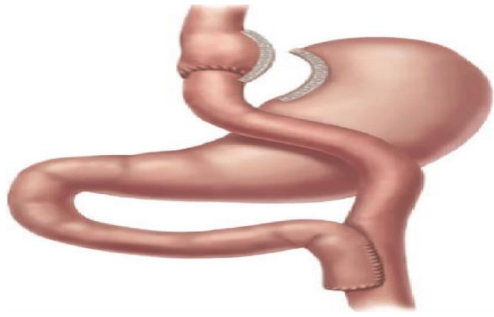


IFSO Bariatric Endoscopy Committee Evidence-Based Review and Position Statement on Endoscopic Sleeve Gastroplasty for Obesity Management

Barham K. Abu Dayyeh^{1,10} · Christine Stier² · Aayed Alqahtani³ · Reem Sharaiha⁴ · Mohit Bandhari⁵ ·
Silvana Perretta⁶ · Singh Pichamol Jirapinyo⁷ · Gerhard Prager⁸ · Ricardo V. Cohen⁹



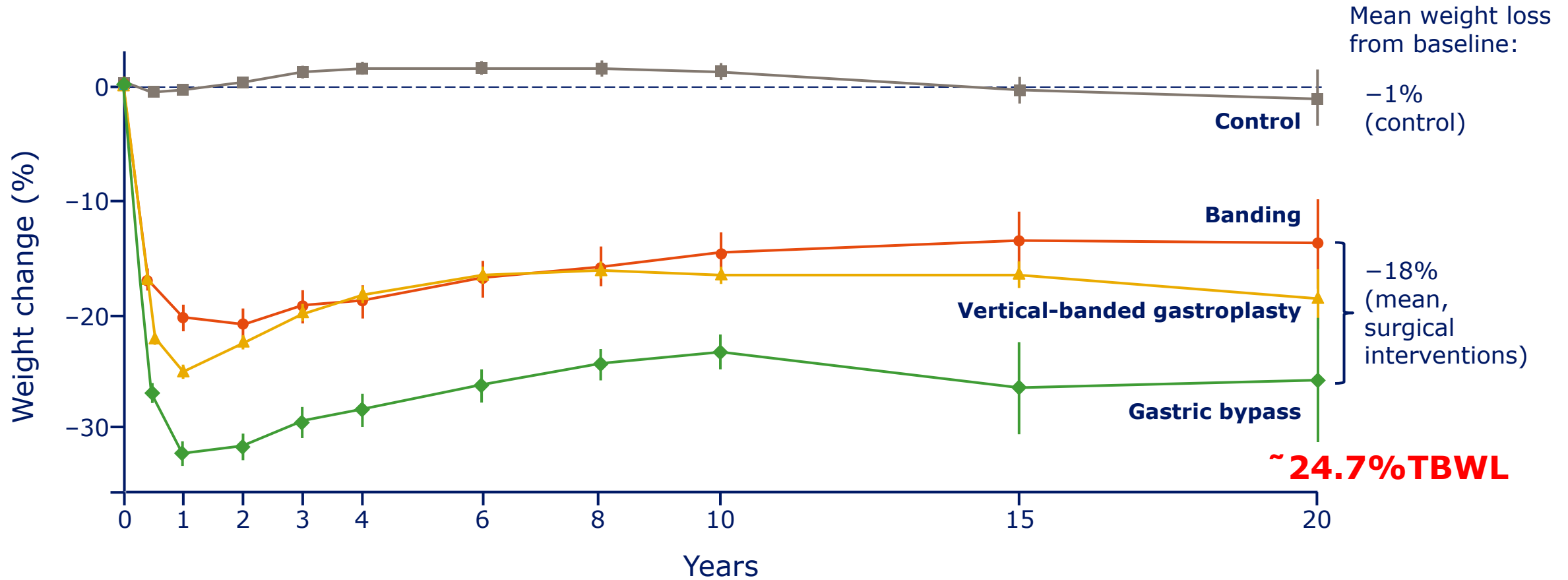




Metabolic and Bariatric Surgery



Long-term significant WL



Data are mean \pm 95% confidence interval

RCTs of metabolic surgery x BMT, at least 2 years of FU, glucocentric outcomes

	Surgical intervention	Follow-up duration, years	Glycaemic target	Proportion reaching glycaemic target (surgical intervention vs current medical treatment), %	Total bodyweight loss (surgical intervention vs current medical treatment), %
Dixon et al ³⁷	AGB	2	FPG <126 mg/dL and HbA _{1c} <6.2% (44.3 mmol/mol), without glucose-lowering agents	73% vs 13%	20% vs 1%
Cohen et al ²¹	RYGB	2	HbA _{1c} <6.5% (47.5 mmol/mol), regardless of glucose-lowering agents	71% vs 51%	26% vs 5%
Simonson et al ³⁸	RYGB	3	FPG <126 mg/dL and HbA _{1c} <6.5% (47.5 mmol/mol) regardless of glucose-lowering agents	42% vs 0%	25% vs 5%
Ikramuddin et al ³⁹	RYGB	5	HbA _{1c} <7% (53.0 mmol/mol), regardless of glucose-lowering agents	55% vs 14%	22% vs 10%
Courcoulas et al ⁴⁰	RYGB vs AGB	5	HbA _{1c} <6.5 (47.5 mmol/mol) or FPG <126 mg/dL, without glucose-lowering agents	30% (RYGB) vs 19% (AGB) vs 0%	25% (RYGB) vs 15% (AGB) vs 6%
Wentworth et al ⁴¹	AGB	5	FPG <126 mg/dL and 2 h blood glucose concentration <200 mg/dL (75 g glucose oral challenge test)	23% vs 9%	12% vs 2%
Schauer et al ⁴²	RYGB vs sleeve gastrectomy	5	HbA _{1c} <6% (42.1 mmol/mol), regardless of glucose-lowering agents	29% (RYGB) vs 23% (sleeve gastrectomy) vs 5%	23% (RYGB) vs 19% (sleeve gastrectomy) vs 5%
Mingrone et al ⁴³	RYGB vs biliopancreatic diversion	10	FPG <100 mg/dL and HbA _{1c} <6.5% (47.5 mmol/mol), without glucose-lowering agents	25% (RYGB) vs 50% (biliopancreatic diversion) vs 5%	37% (RYGB) vs 42% (biliopancreatic diversion) vs 7%

HbA_{1c}=glycated haemoglobin. FPG=fasting plasma glucose. AGB=adjustable gastric banding. RYGB=Roux-en-Y gastric bypass.

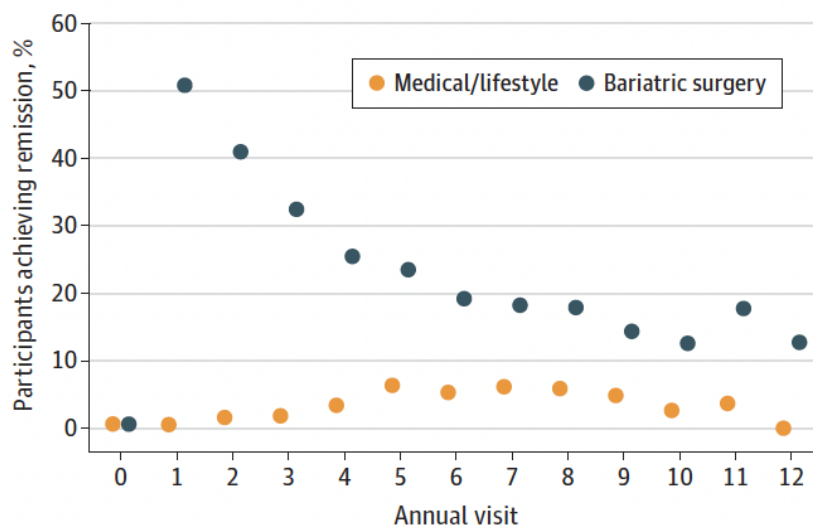
Table 1: Randomised controlled trials with follow-up duration of at least 2 years comparing bariatric surgery with current medical treatment

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

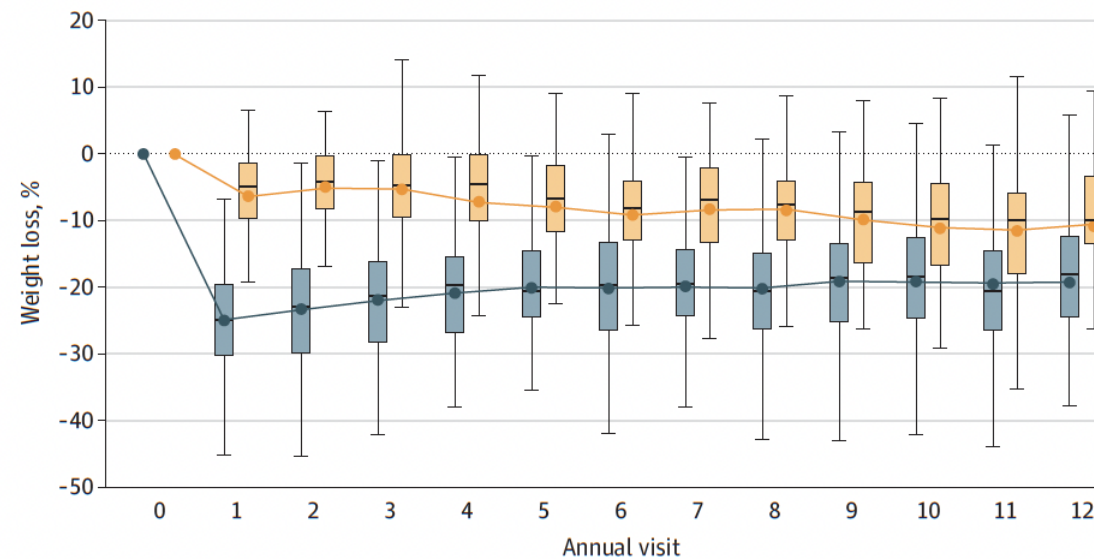
Figure 3. Diabetes Remission



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

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

C Weight loss



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

The image is a composite of two parts. The left side features a 3D anatomical illustration of a kidney, showing its reddish-brown outer cortex and the internal renal pyramids and pelvis. The right side is a photograph of a hand holding a stethoscope against a person's arm, with a blurred background showing a blue medical bag with a white cross symbol. The text "Non- glucocentric endpoints" is centered across the middle of the image in a white, sans-serif font.

Non- glucocentric endpoints

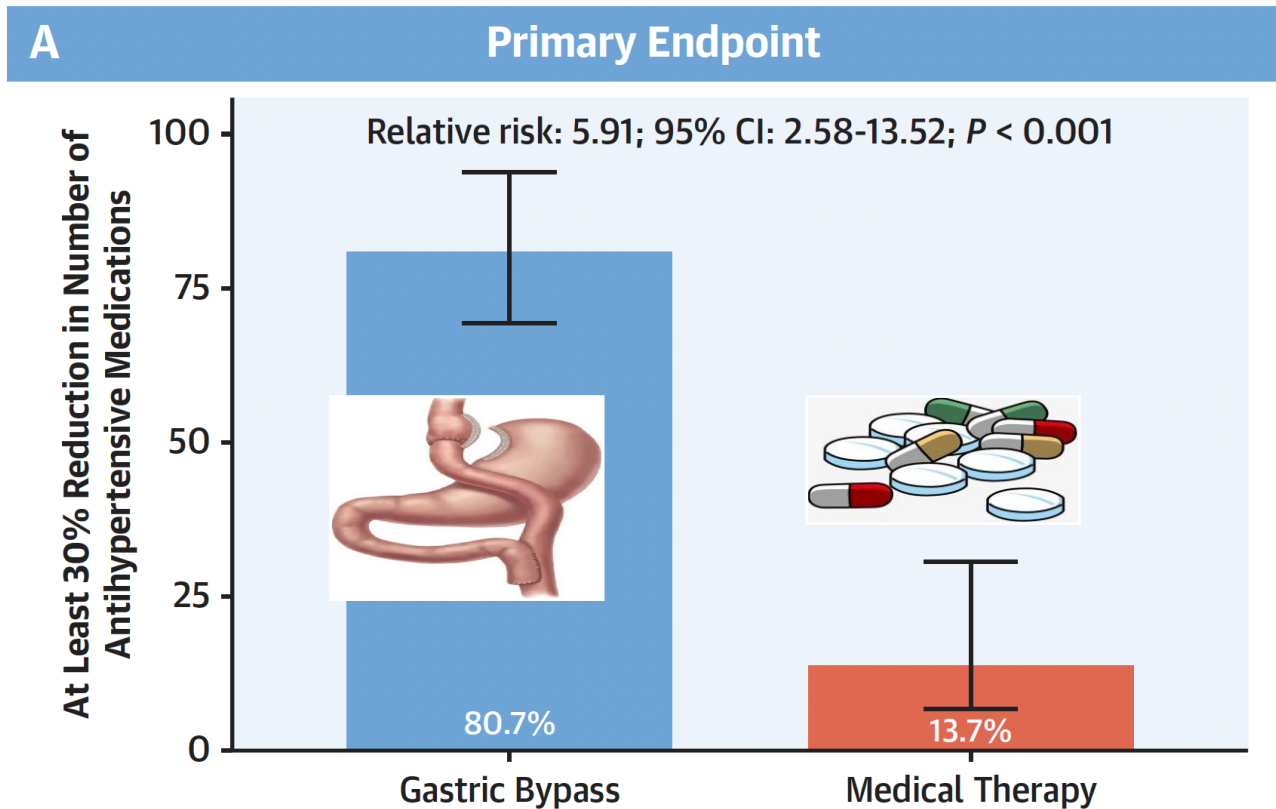
Randomized Trial of Effect of Bariatric Surgery on Blood Pressure After 5 Years



Carlos A. Schiavon, MD,^{a,b} Alexandre B. Cavalcanti, MD,^a Juliana D. Oliveira, CN,^{a,b} Rachel H.V. Machado, CN,^a Eliana V. Santucci, Pr,^a Renato N. Santos, STAT,^a Julia S. Oliveira, STAT,^a Lucas P. Damiani, STAT,^a Débora Junqueira, MD,^a Helio Halpern, MD,^c Frederico de L.J. Monteiro, MD,^c Patricia M. Noujaim, MD,^b Ricardo V. Cohen, MD,^d Marcio G. de Sousa, MD,^e Luiz A. Bortolotto, MD,^f Otavio Berwanger, MD,^g Luciano F. Drager, MD^{f,h,i}



Primary outcome was at least a 30% reduction in total number of antihypertensive medications

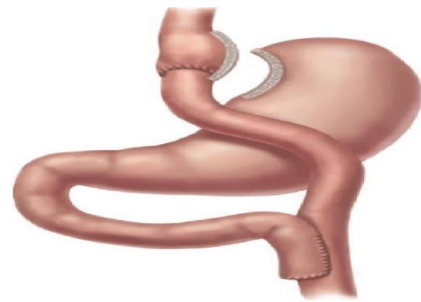


MOMS TRIAL

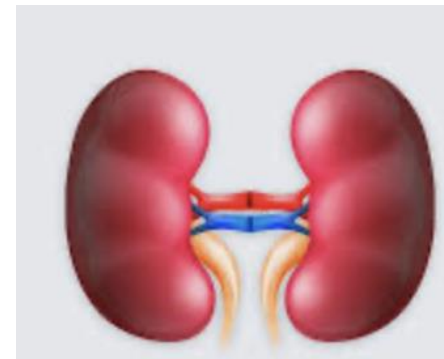
Gastric bypass *versus* best medical treatment for diabetic kidney disease: 5 years follow up of a single-centre open label randomised controlled trial

Ricardo V. Cohen,^{a,*} Tiago Veiga Pereira,^{b,c} Cristina Mamédio Aboud,^a Tarissa Beatrice Zanata Petry,^a José Luis Lopes Correa,^a Carlos Aurélio Schiavon,^d Carlos Eduardo Pompílio,^a Fernando Nogueira Quirino Pechy,^a Ana Carolina Calmon da Costa Silva,^a Lívia Porto Cunha da Silveira,^a Pedro Paulo de Paris Caravatto,^a Helio Halpern,^a Frederico de Lima Jacy Monteiro,^a Bruno da Costa Martins,^a Rogerio Kuga,^a Thais Mantovani Sarian Palumbo,^a Allon N. Friedman,^e and Carel W. le Roux^{f,g}

The Lancet , online Nov 11,2022



X



MOMS trial

Endpoints

- **Primary endpoint**

- uACR < 30 mg/g

- **Secondary endpoints**

- CKD remission

- Metabolic control (A1c < 6%; FPG < 100 mg/dl; LDL < 100 mg/dl (< 70 if CV+); HDL > 50; TG < 150 mg/dl; SBP < 130 mmHg ; DBP < 80 mmHg

- Weight-loss

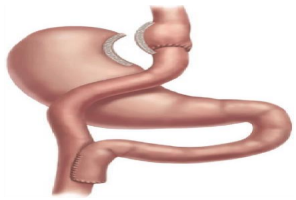
- Use of T2D medications

- Neuropathy/Retinopathy

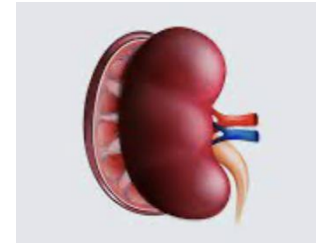
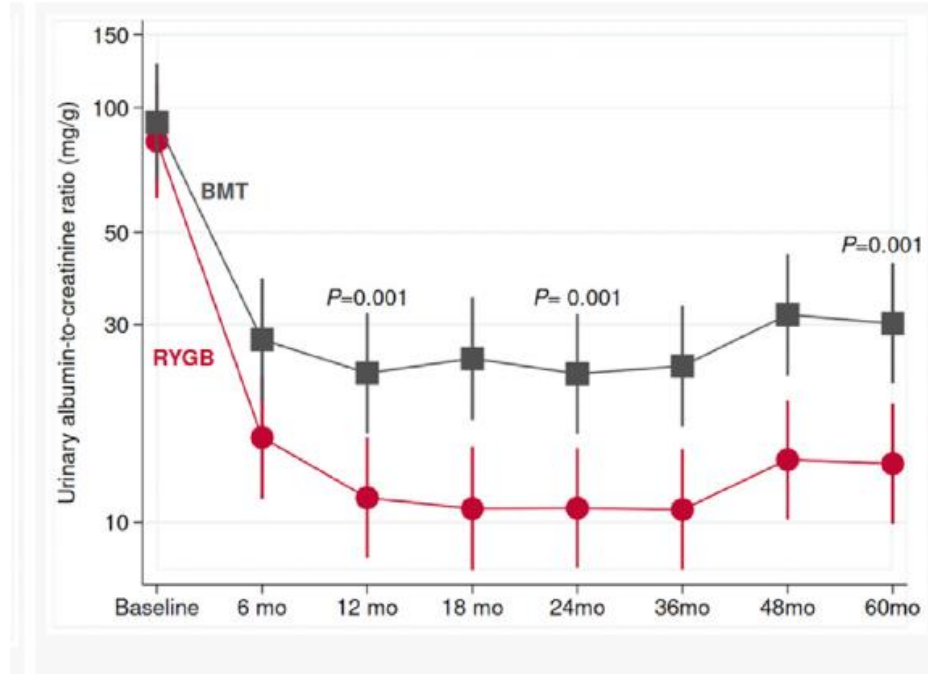
- QOL

MOMS trial – 5 years outcomes

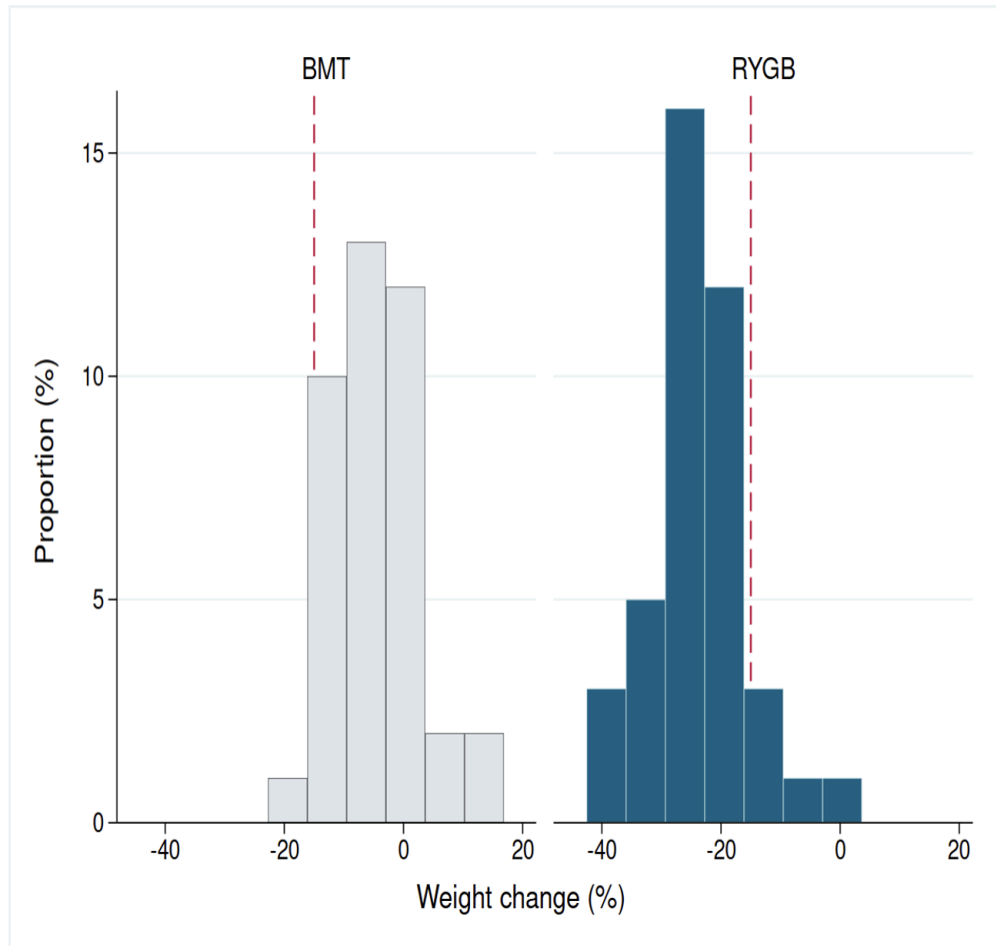
Primary outcome – uACR- continuous variable
B



+ BMT



*The geometric mean for albuminuria levels
was 46%
lower after RYGB (P = 0.001)*

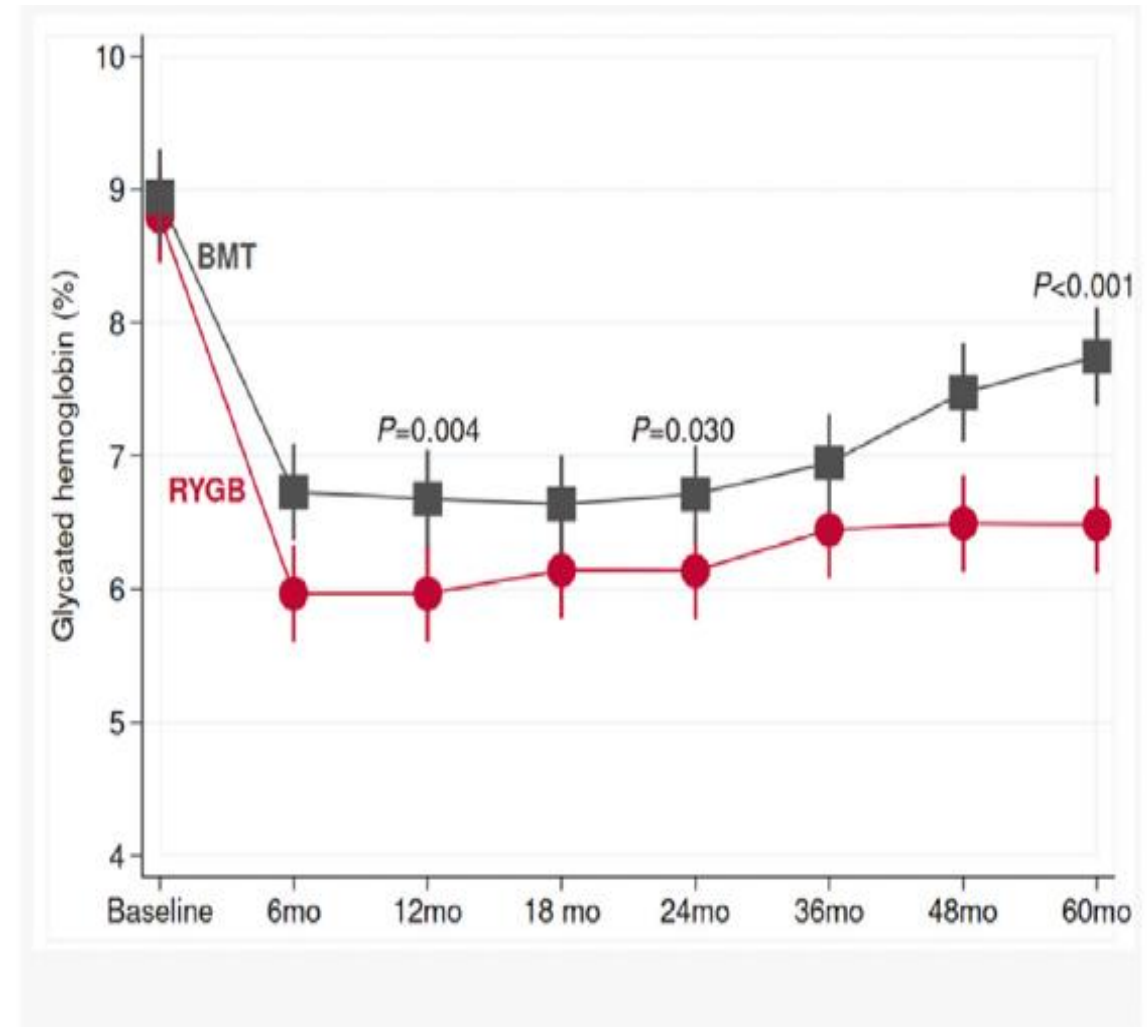


Only 22.5% BMT achieved >15% TBWL

90% RYGB >15% TBWL

NO BMT pt reached normal BMI

53% after RYGB ($P < 0.001$)



Glycemic control

Cv events and mortality

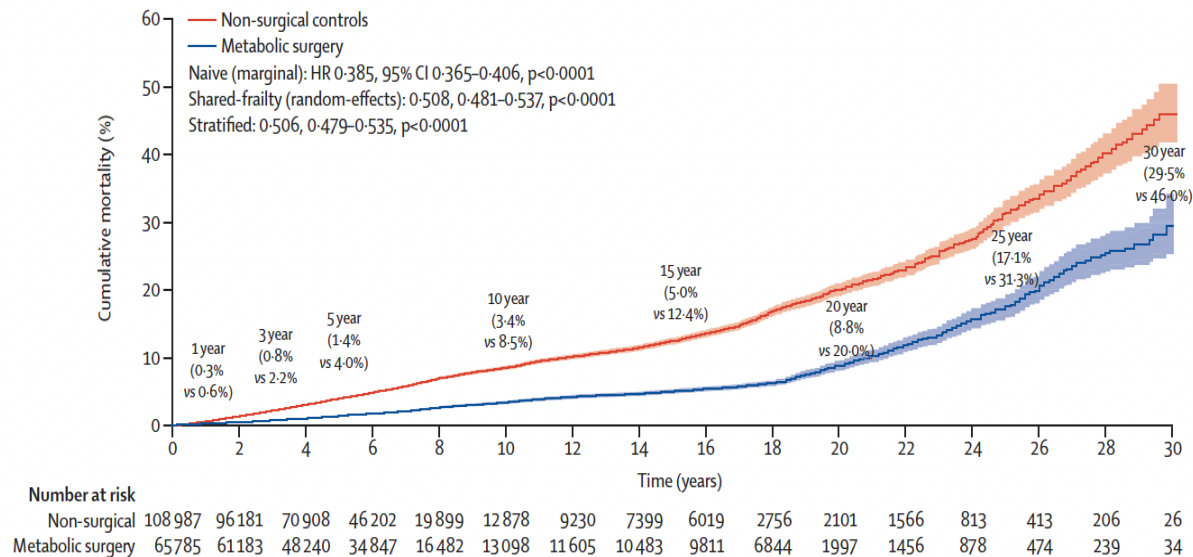


Association of metabolic-bariatric surgery with long-term survival in adults with and without diabetes: a one-stage meta-analysis of matched cohort and prospective controlled studies with 174 772 participants



Lancet, 05/21

Nicholas L Syn*, David E Cummings*, Louis Z Wang*, Daryl J Lin*, Joseph J Zhao, Marie Loh, Zong Jie Koh, Claire Alexandra Chew, Ying Ern Loo, Bee Choo Tai, Guowei Kim, Jimmy Bok-Yan So, Lee M Kaplan, John B Dixon, Asim Shabbir



✓ Reduction of all-cause mortality

✓ Overall ~ 50%

✓ Pre-existing T2D ~ 60% reduction

✓ No T2D @ baseline ~ 30% reduction

Metanalysis

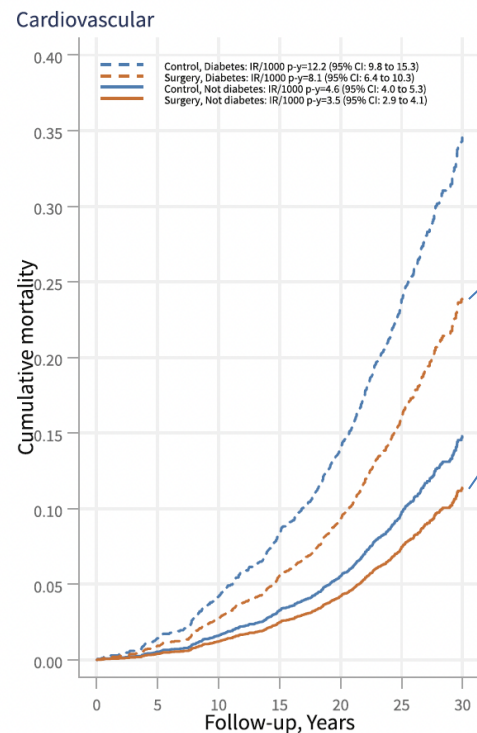
174.772 pts

Life expectancy after bariatric surgery or usual care in patients with or without baseline type 2 diabetes in Swedish Obese Subjects

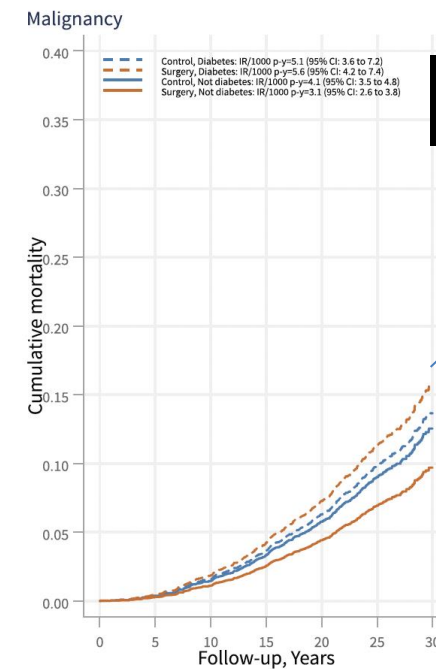
July 12, 2023

Lena M. S. Carlsson¹, Björn Carlsson^{1,2}, Peter Jacobson¹, Cecilia Karlsson^{1,3}, Johanna C. Andersson-Assarsson¹, Felipe M. Kristensson^{1,4}, Sofie Ahlin^{1,5}, Per-Arne Svensson^{1,6}, Magdalena Taube¹, Ingmar Näslund⁷, Kristjan Karason¹, Markku Peltonen⁸ and Kajsa Sjöholm¹✉

26 years FU, SOS study

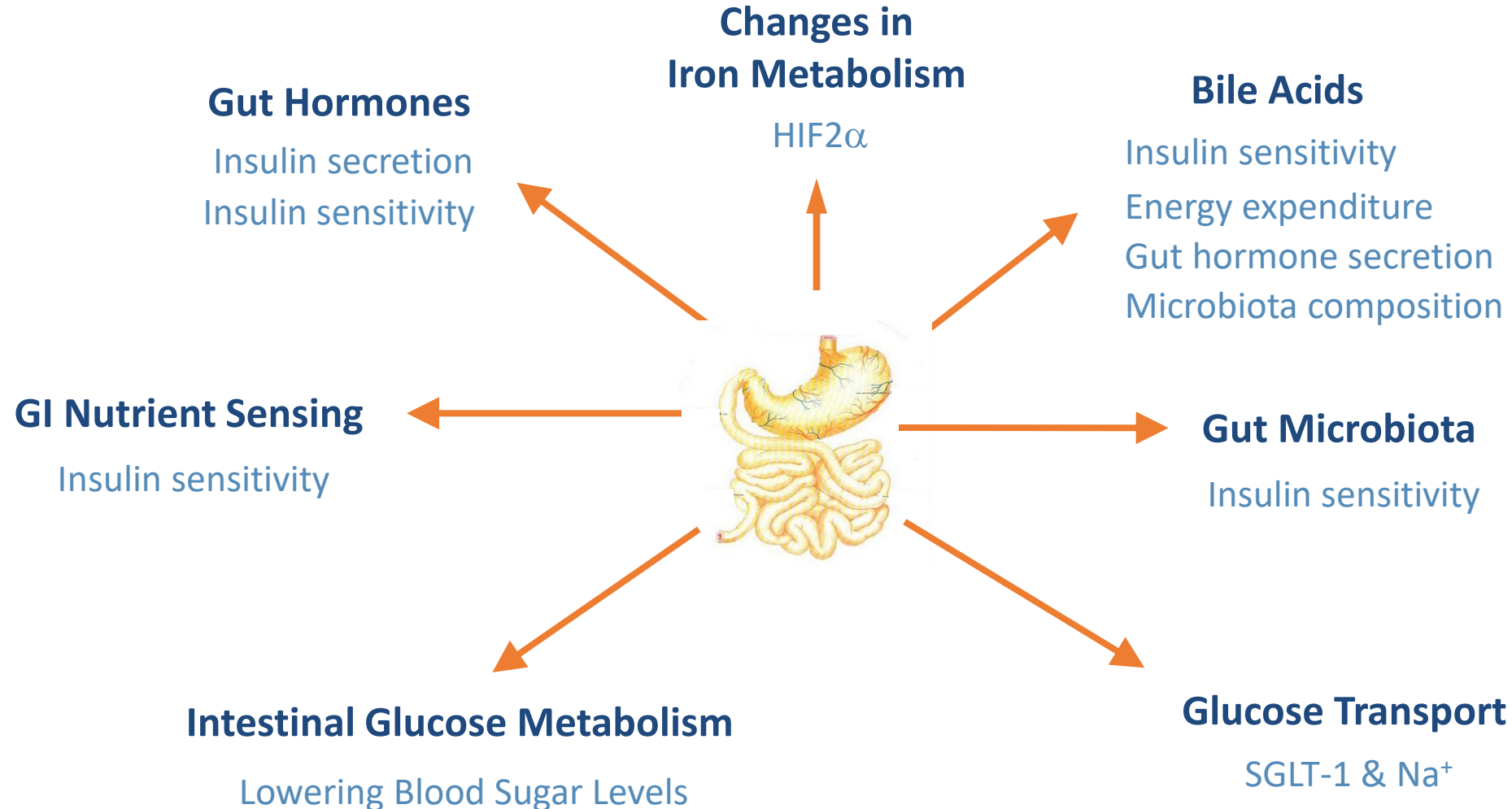


Less fatal CVE, w or w/o T2D



Less cancer deaths, w or w/o T2D

MBS physiological Mechanisms



Complications and mortality continuous decrease

Campos et al

Annals of Surgery • Volume 271, Number 2, February 2020

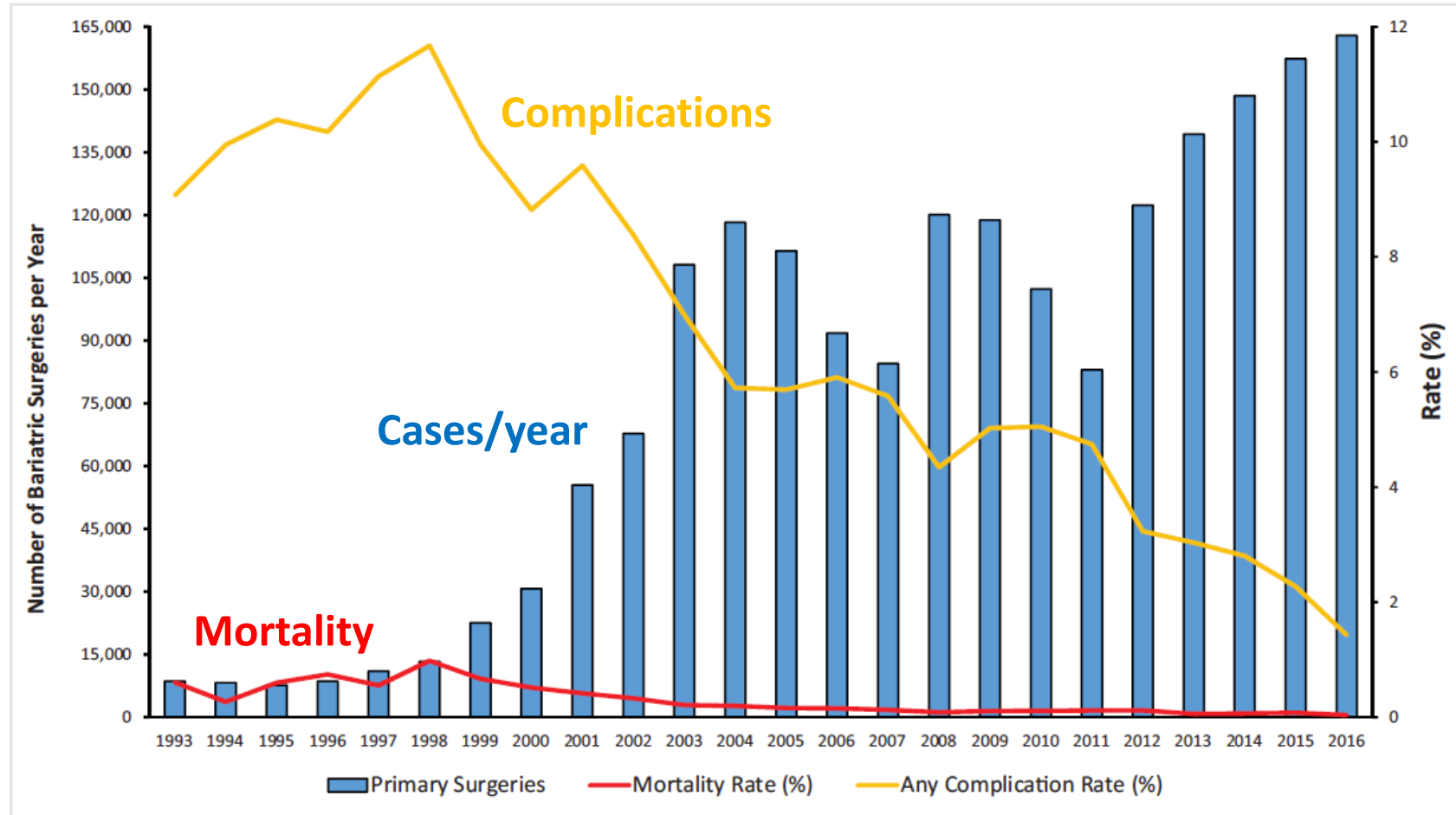


FIGURE 2. Number of inpatient primary bariatric surgery procedures and initial admission complication and mortality rates in the United States from 1993 to 2016.

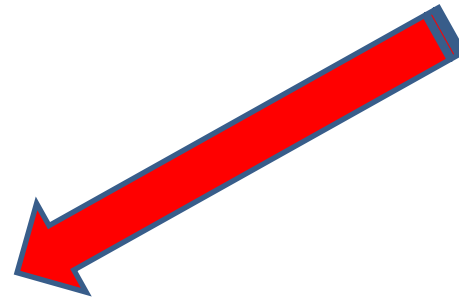
There is no magic bullet for obesity



*Ildiko Lingvay, Priya Sumithran,
Carel W le Roux, *Ricardo V Cohen*
ricardo.cohen@haoc.com.br

Lancet DE, July 2023

**DECISIONS ARE MORE IMPORTANT
THAN INCISIONS**



BEST OUTCOMES

OBESITY TREATMENT

An infographic titled "OBESITY TREATMENT" showing four paths leading to a central goal. The paths are labeled "MEDICATIONS", "ENDOSCOPIC SLEEVE GASTROPLASTY", "BARIATRIC SURGERY", and "COMBINATION". Below these paths is a wide base labeled "LIFESTYLE INTERVENTIONS". Silhouettes of people are walking on the paths. The background is a light yellow sky with clouds.

MEDICATIONS

ENDOSCOPIC
SLEEVE
GASTROPLASTY

BARIATRIC
SURGERY

COMBINATION

LIFESTYLE INTERVENTIONS



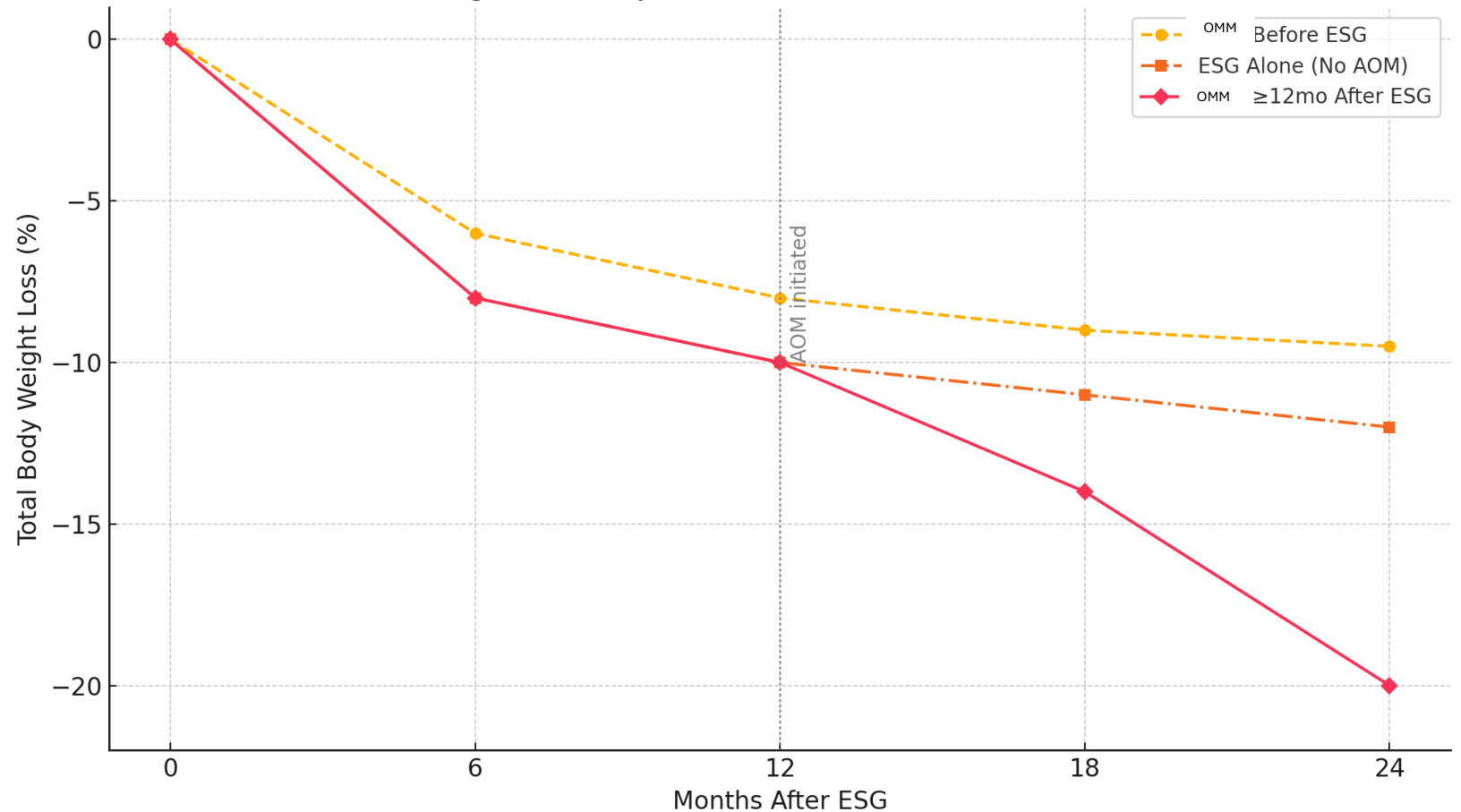
Outcomes of concomitant antiobesity medication use with endoscopic sleeve gastroplasty in clinical US settings

Khushboo Gala^a, Wissam Ghusn^a, Vitor Brunaldi^a, Christopher McGowan^b, Reem Z. Sharaiha^c, Daniel Maselli^b, Brandon Vanderwel^d, Prashant Kedia^e, Michael Ujiki^f, Eric Wilson^g, Eric J. Vargas^a, Andrew C. Storm^a, Barham K. Abu Dayyeh^{a,*}

^aNorthwestern University, Chicago, IL, USA; ^bUniversity of Michigan, Ann Arbor, MI, USA; ^cUniversity of California, San Diego, CA, USA; ^dUniversity of Texas, Houston, TX, USA; ^eUniversity of Illinois, Chicago, IL, USA; ^fUniversity of Michigan, Ann Arbor, MI, USA; ^gUniversity of Michigan, Ann Arbor, MI, USA

70% Sema and Liraglutide
30% other agents, including NB

Weight Loss Trajectories with ESG and OMM Timing

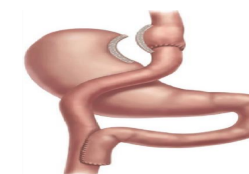




Bariatric Surgery: There Is a Room for Improvement to Reduce Mortality in Patients with Type 2 Diabetes

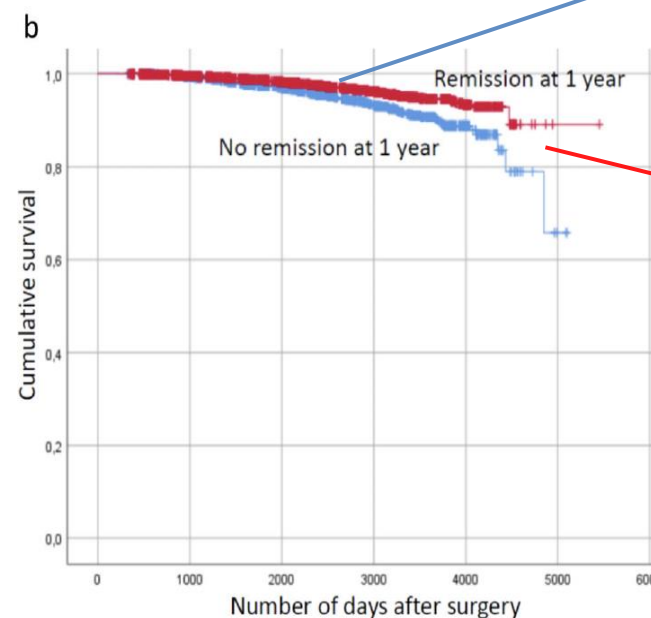
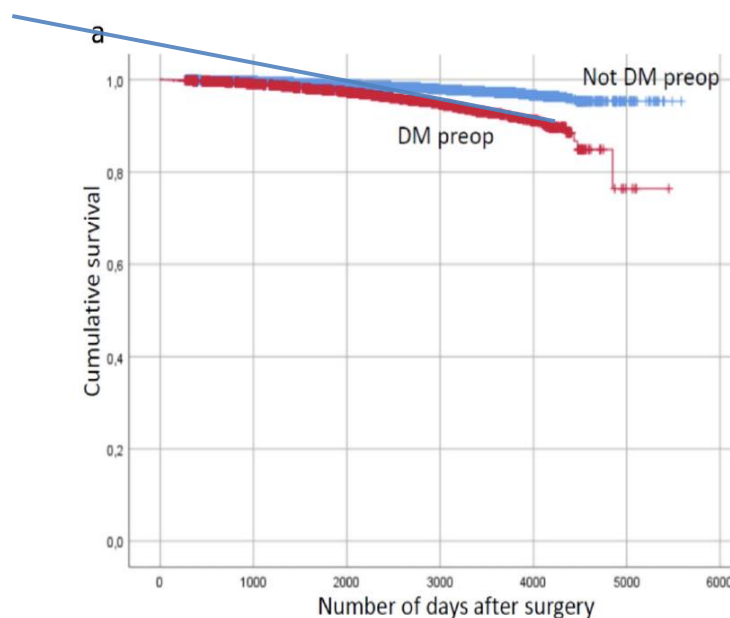
Carel W. le Roux¹ · Johan Ottosson^{2,3} · Erik Näslund^{2,4} · Ricardo V. Cohen⁵ · Erik Stenberg^{2,3} · Magnus Sundbom^{2,6} · Ingmar Näslund^{2,3}

Received: 6 July 2020 / Revised: 13 August 2020 / Accepted: 14 August 2020
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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

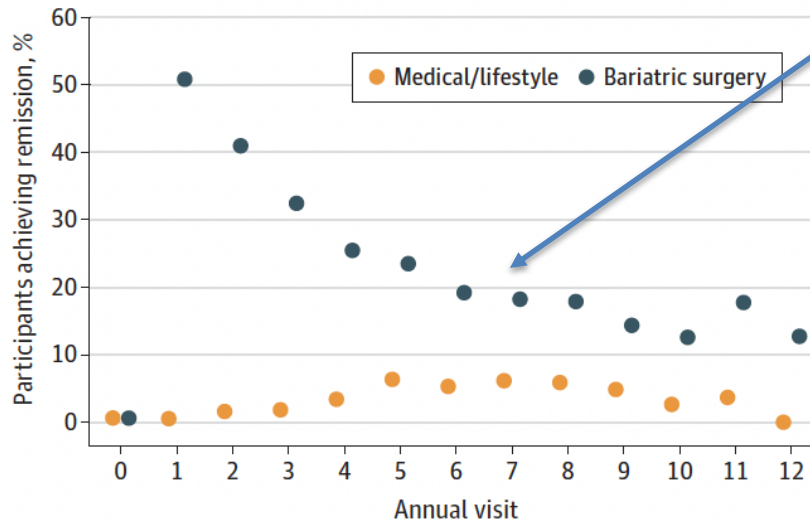
Long-Term Outcomes of Medical Management vs Bariatric Surgery in Type 2 Diabetes

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

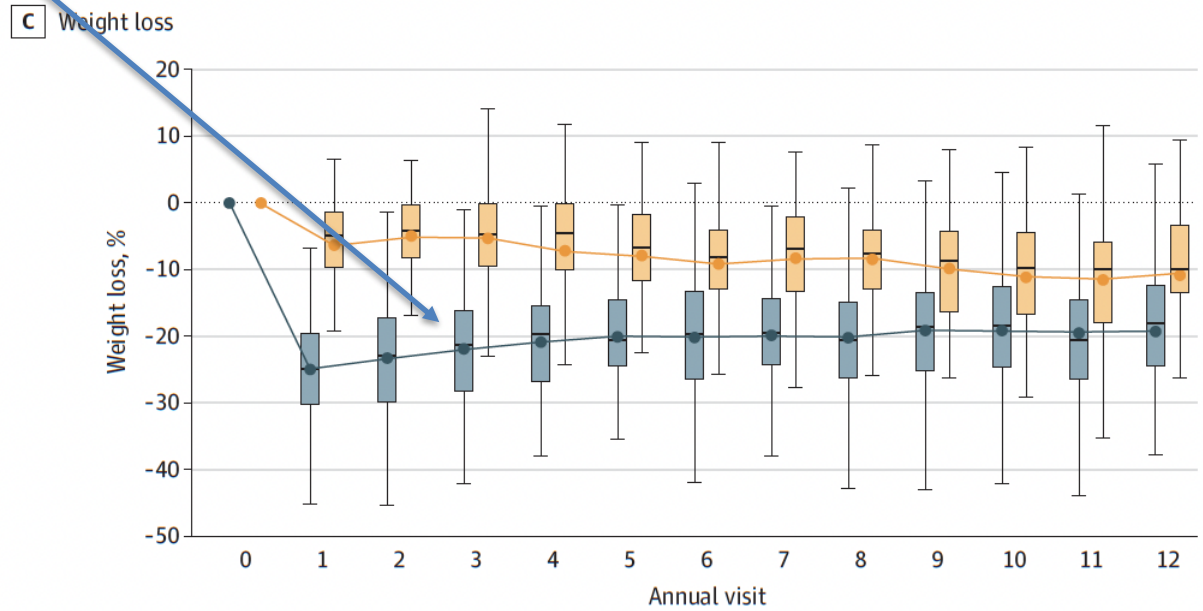
2024

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

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

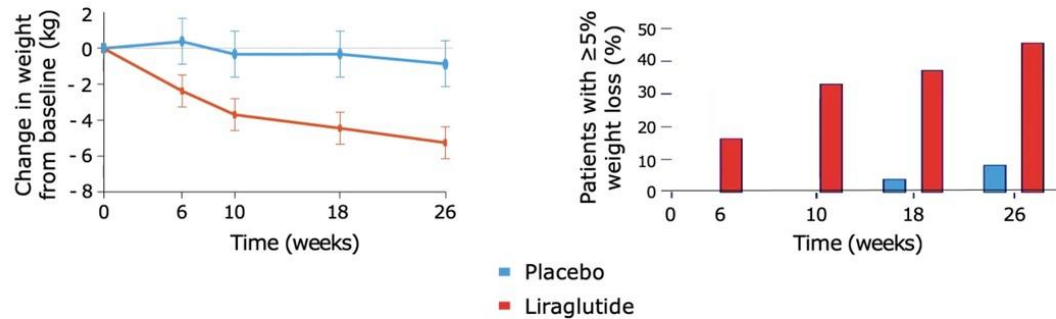
Mok et al, 2023

The BARI-OPTIMISE Randomized Clinical Trial

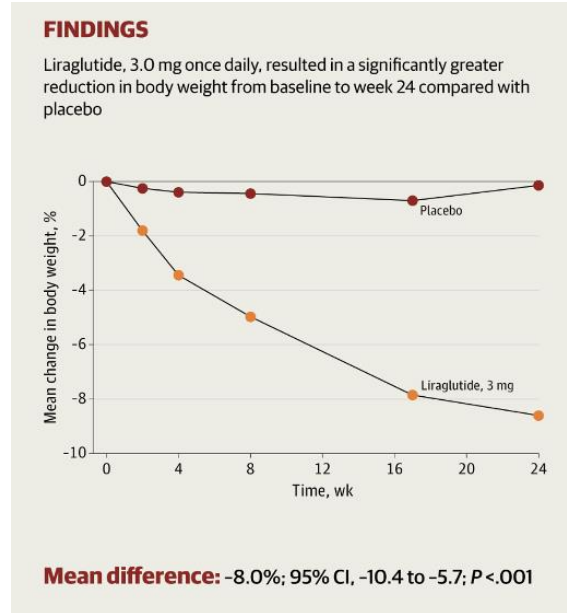
Postoperative pharmacotherapy augments surgical weight loss

GRAVITAS Study

Liraglutide after Gastric Bypass in T2D



Miras AD et al., Lancet Diabetes Endocrinol 2019



Cohen RV et al, BJS, 2024

OMMs after MBS

	Grade	Consensus (%)	Nr.of rounds	Nr.of total votes
When used after MBS, there appears to be no increased incidence of side effects of OMMs compared to non-surgical patients	A	97	3	34

Side Effects Reported with Liraglutide post-MBS in BARI-OPTIMISE compared with SCALE

Adverse Events (AE)- in the BARI-OPTIMISE Study Population

Event	Participants who experienced an AE, No. (%)		
	Placebo (n = 35)	Liraglutide (n = 35)	Total (N = 70)
Total	20 (57)	28 (80)	48 (67)
Total AEs, No. ^a	75	37	112
Gastrointestinal events			
Nausea	7 (20)	18 (51)	25 (36)
Diarrhea	2 (6)	2 (6)	4 (6)
Constipation	2 (6)	9 (26)	11 (16)
Vomiting	1 (3)	1 (3)	2 (3)
Abdominal pain	1 (3)	2 (6)	3 (4)
Abdominal bloating	0	1 (3)	1 (1)
Dyspepsia	0	1 (3)	1 (1)

Table 3. Adverse Events and Serious Adverse Events.^a

Event	Liraglutide (N=2481)			Placebo (N=1242)		
	No. of Patients (%)	No. of Events	Event Rate per 100 Exposure-Years	No. of Patients (%)	No. of Events	Event Rate per 100 Exposure-Years
Adverse events in ≥5% of patients	1992 (80.3)	7191	321.8	786 (63.3)	2068	193.7
Nausea	997 (40.2)	1429	63.9	183 (14.7)	223	20.9
Diarrhea	518 (20.9)	754	33.7	115 (9.3)	142	13.3
Constipation	495 (20.0)	593	26.5	108 (8.7)	121	11.3
Vomiting	404 (16.3)	597	26.7	51 (4.1)	62	5.8
Dyspepsia	236 (9.5)	282	12.6	39 (3.1)	44	4.1
Upper abdominal pain	141 (5.7)	171	7.7	43 (3.5)	49	4.6
Abdominal pain	130 (5.2)	163	7.3	43 (3.5)	53	5.0
Nasopharyngitis	427 (17.2)	586	26.2	234 (18.8)	302	28.3
Upper respiratory tract infection	213 (8.6)	247	11.1	122 (9.8)	149	14.0
Sinusitis	128 (5.2)	141	6.3	73 (5.9)	95	8.9
Influenza	144 (5.8)	170	7.6	66 (5.3)	84	7.9
Headache	327 (13.2)	443	19.7	154 (12.4)	220	20.6
Dizziness	167 (6.7)	203	9.1	60 (4.8)	65	6.1
Decreased appetite	267 (10.8)	283	12.7	38 (3.1)	39	3.7
Back pain	171 (6.9)	210	9.4	105 (8.5)	121	11.3
Arthralgia	125 (5.0)	133	6.0	71 (5.7)	80	7.5
Fatigue	185 (7.5)	203	9.1	65 (5.2)	72	6.7
Injection-site hematoma	142 (5.7)	154	6.9	93 (7.5)	101	9.5

Mok et al. JAMA Surg 2023 (BARI-OPTIMISE); SCALE. Pi-Sunyer NEJM 2015

Side Effects Reported with Liraglutide post-MBS: Retrospective study of 117 patients compared to SCALE

TABLE 2 Side effects for liraglutide 3.0 mg

Symptom	n (%) ^a
Nausea	34 (29.1%)
Constipation	13 (11.1%)
Diarrhoea	8 (6.8%)
Fatigue	7 (6.0%)
Headache	4 (3.4%)
Rash	4 (3.4%)
Indigestion	3 (2.6%)
Vomiting	3 (2.6%)
Dry mouth	3 (2.6%)
Bloating	2 (1.7%)
Sweating	2 (1.7%)
Other ^b	9 (7.7%)

^aPercent calculated as [(number of patients reporting the side effect)/117] × 100.

^bOther includes abdominal pain (n = 1), bruising (n = 1), decreased glomerular filtration rate (n = 1), depression (n = 1), flu-like symptoms (n = 1), heartburn (n = 1), hot flashes (n = 1), gas (n = 1) and pancreatitis (n = 1).

Table 3. Adverse Events and Serious Adverse Events.^a

Event	Liraglutide (N=2481)			Placebo (N=1242)		
	No. of Patients (%)	No. of Events	Event Rate per 100 Exposure-Years	No. of Patients (%)	No. of Events	Event Rate per 100 Exposure-Years
Adverse events in ≥5% of patients	1992 (80.3)	7191	321.8	786 (63.3)	2068	193.7
Nausea	997 (40.2)	1429	63.9	183 (14.7)	223	20.9
Diarrhea	518 (20.9)	754	33.7	115 (9.3)	142	13.3
Constipation	495 (20.0)	593	26.5	108 (8.7)	121	11.3
Vomiting	404 (16.3)	597	26.7	51 (4.1)	62	5.8
Dyspepsia	236 (9.5)	282	12.6	39 (3.1)	44	4.1
Upper abdominal pain	141 (5.7)	171	7.7	43 (3.5)	49	4.6
Abdominal pain	130 (5.2)	163	7.3	43 (3.5)	53	5.0
Nasopharyngitis	427 (17.2)	586	26.2	234 (18.8)	302	28.3
Upper respiratory tract infection	213 (8.6)	247	11.1	122 (9.8)	149	14.0
Sinusitis	128 (5.2)	141	6.3	73 (5.9)	95	8.9
Influenza	144 (5.8)	170	7.6	66 (5.3)	84	7.9
Headache	327 (13.2)	441	19.7	154 (12.4)	220	20.6
Dizziness	167 (6.7)	203	9.1	60 (4.8)	65	6.1
Decreased appetite	267 (10.8)	283	12.7	38 (3.1)	39	3.7
Back pain	171 (6.9)	210	9.4	105 (8.5)	121	11.3
Arthralgia	125 (5.0)	133	6.0	71 (5.7)	80	7.5
Fatigue	185 (7.5)	203	9.1	65 (5.2)	72	6.7
Injection-site hematoma	142 (5.7)	154	6.9	93 (7.5)	101	9.5

Wharton S. et al. Clinical Obesity. 2019;9:e12323; SCALE. Pi-Sunyer NEJM 2015

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)

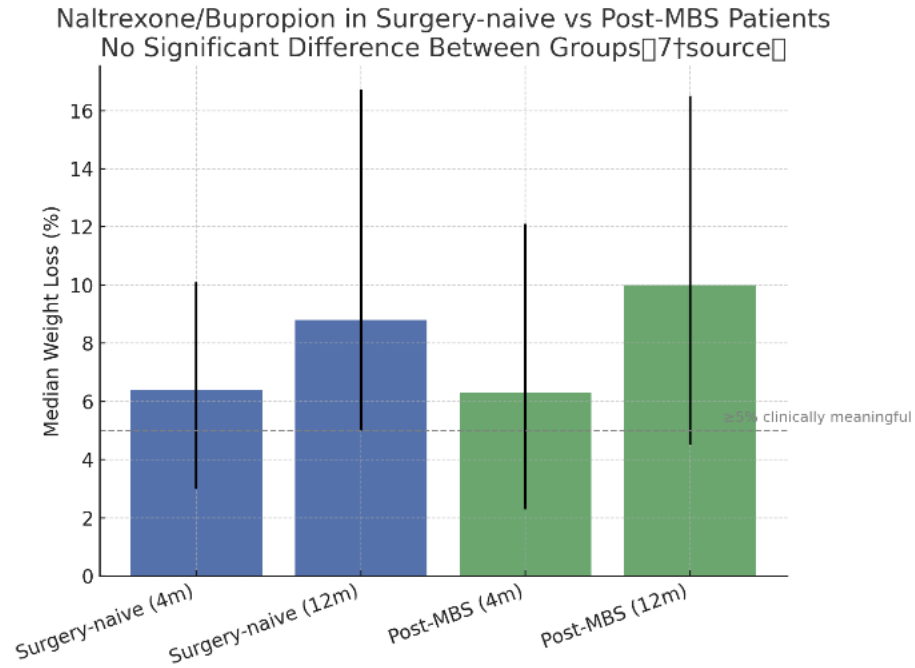
Treatment Protocol & Dosing

- **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

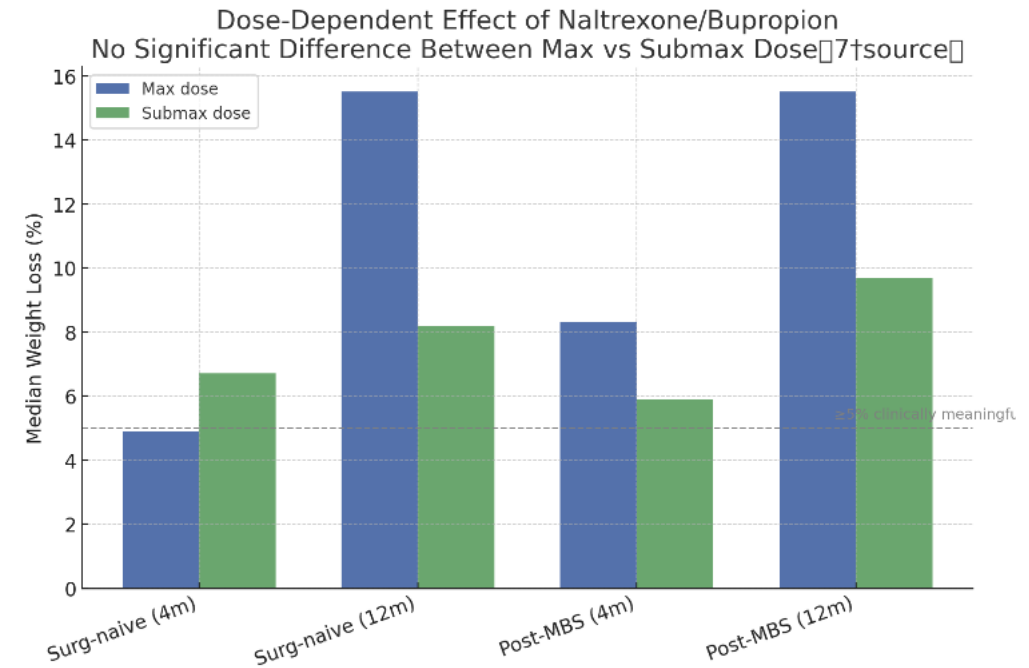
	Week 1	Week 2	Week 3	Week 4 and Beyond
 AM: Take with breakfast	 1 tablet	 1 tablet	 2 tablets	 2 tablets
 PM: Take before dinner		 1 tablet	 1 tablet	 2 tablets

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...

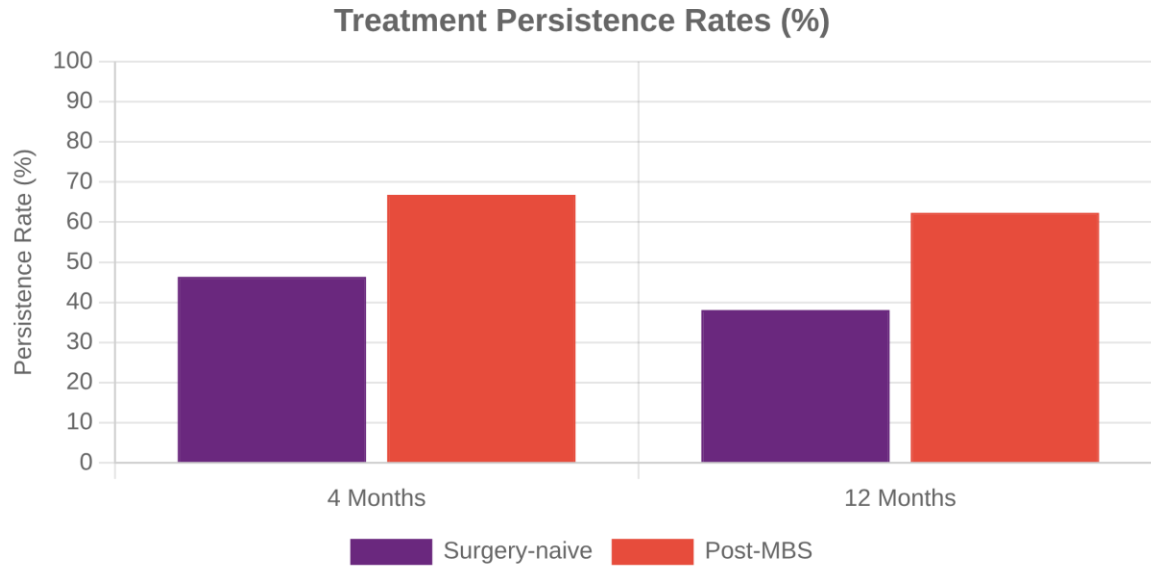
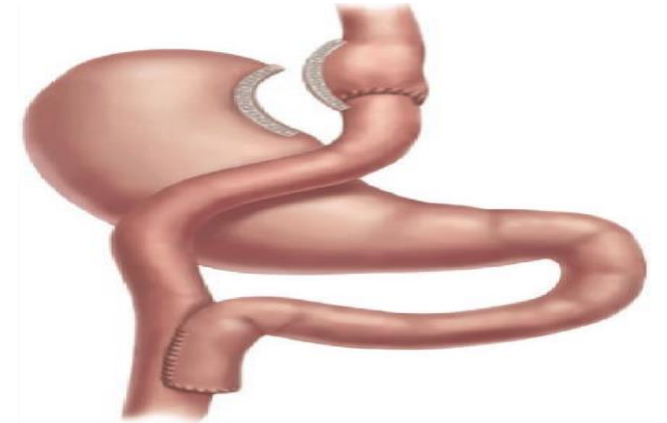


No dose-dependent effect

● **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 or 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



REVIEW



Medium and Long-Term Weight Loss After Revisional Bariatric Surgery: A Systematic Review and Meta-Analysis

Eduardo L. S. Bastos¹ · Wilson Salgado Jr.² · Anna C. B. Dantas³ · Tiago R. Onzi⁴ · Lyz B. Silva⁵ · Álvaro Albano⁶ · Luca S. Tristão⁷ · Clara L. dos Santos⁷ · Antonio Silvinato⁸ · Wanderley M. Bernardo⁹ · For the Scientific Committee of the Brazilian Society of Bariatric and Metabolic Surgery (SBCBM)

Table 2 Analysis of the quality of evidence (GRADE) in relation to the overall rate of occurrence of the assessed outcomes

Certainty assessment								Patients (n)	Certainty of evidence
Outcome	Studies (n)	Study design	Risk of bias	Inconsistency	Indirectness	Imprecision	Others considerations		
BMI	23	Observational	NS	VS ($I^2 > 75$)	NS	NS	None	1,602	⊕○○○ Very low
%EWL	18	Observational	NS	VS ($I^2 > 75$)	NS	NS	Publication Bias Strongly Suspected	1,031	⊕○○○ Very low
%EBMIL	7	Observational	NS	VS ($I^2 > 75$)	NS	NS	None	350	⊕○○○ Very low
%TWL	16	Observational	NS	VS ($I^2 > 75$)	NS	NS	None	888	⊕○○○ Very low

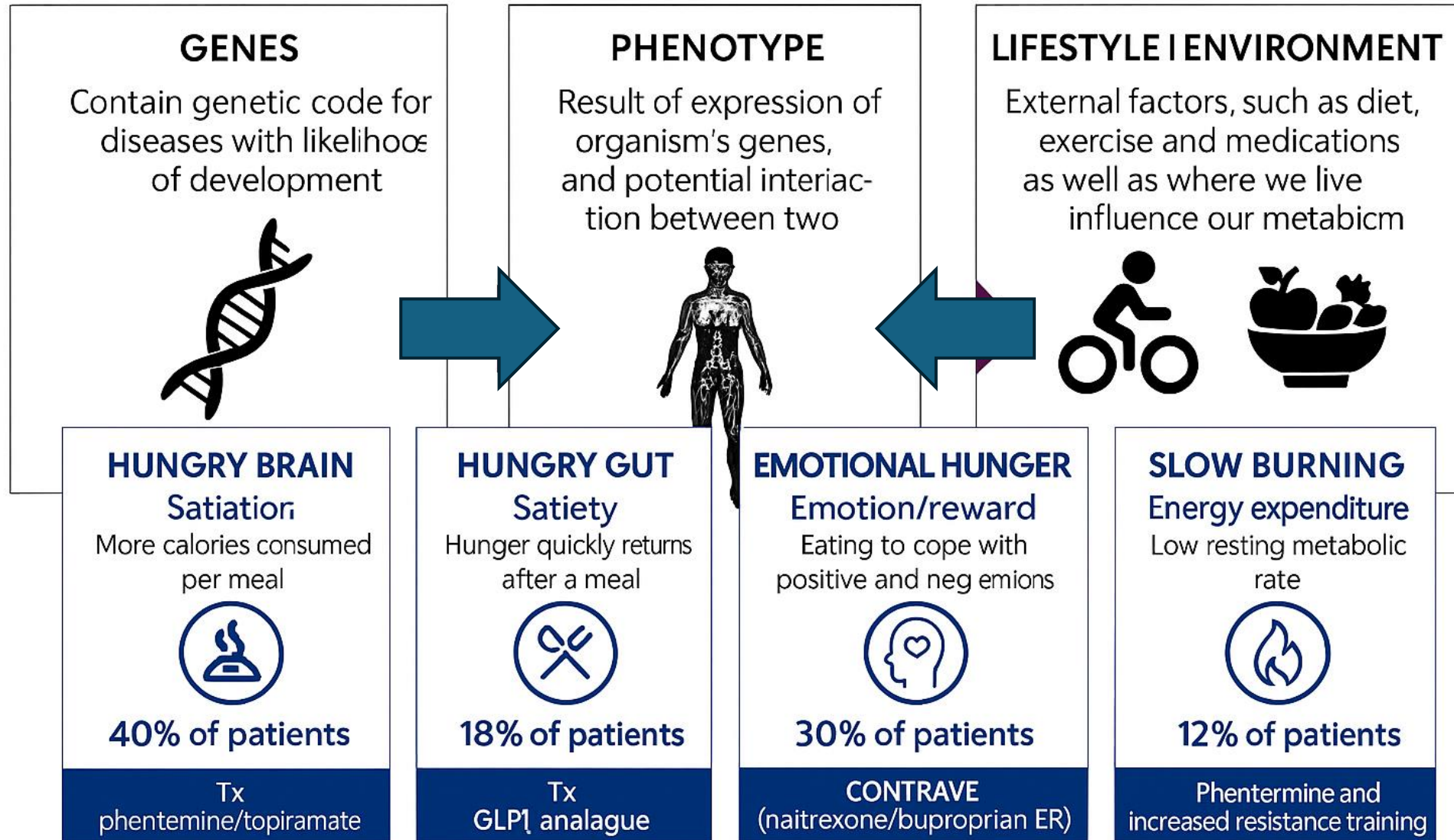
NS, not serious; VS, very serious

OMMs **after** MBS

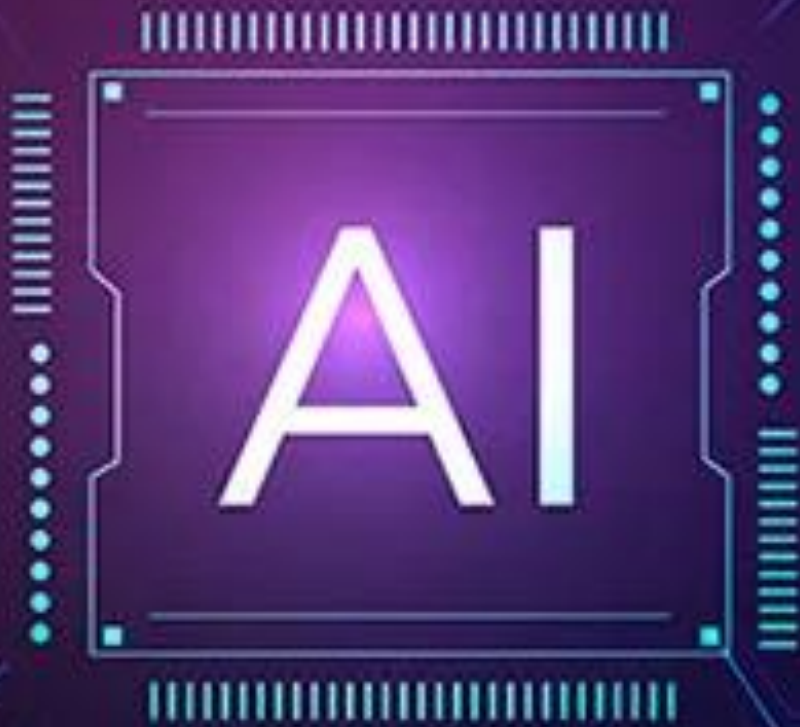
	Grade	Consensus (%)	Nr.of rounds	Nr.of total votes
Future research is needed to identify predictors of which patients are likely to derive substantial benefit from combined pharmacosurgical therapy for obesity and its complications	A+	100	3	35

- ✓ Precision Medicine
- ✓ Phenotyping persons with obesity

Precision medicine in obesity treatment



Personalized Treatment Selection



AI Analyzes medical history, exams and past treatments to recommend which patients **will respond best to surgery, pharmacotherapy, combined strategies and which OMM**

AI Clinical Risk Stratification and phenotyping

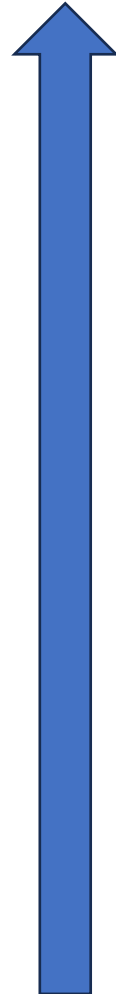
Obesity treatment TODAY

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

ESG in selected pts

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 multiple wings and a central courtyard is prominent. The text "Thank You" is overlaid on a yellow rectangular background in the upper center of the image.

Thank You

ricardo.cohen@haoc.com.br

X= @rvcohen

Disclosures

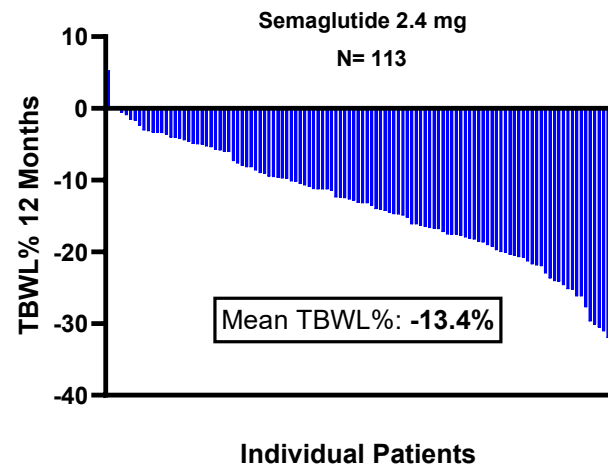
Gila Therapeutics and Phenomix Sciences have licensed Dr. Acosta's research technologies from University of Florida and Mayo Clinic.

Consultant Fees in the last 5 years from Structure Therapeutics, Rhythm Pharmaceuticals, Gila Therapeutics, Amgen, General Mills, Regeneron, Boehringer Ingelheim, Novo Nordisk, Currax, Nestle, Phenomix Sciences, Busch Health, RareDiseases.

Funding support from the National Institute of Health, Delaney Foundation, Dairy Management Institute, Vivus Pharmaceuticals, Apollo Endosurgery, Satiogen Pharmaceuticals, Spatz Medical, Rhythm Pharmaceuticals, Regeneron, Boehringer Ingelheim, Novo Nordisk.

Obstacles In the obesity pandemic

HETEROGENEOUS OUTCOMES¹



HIGH COST²

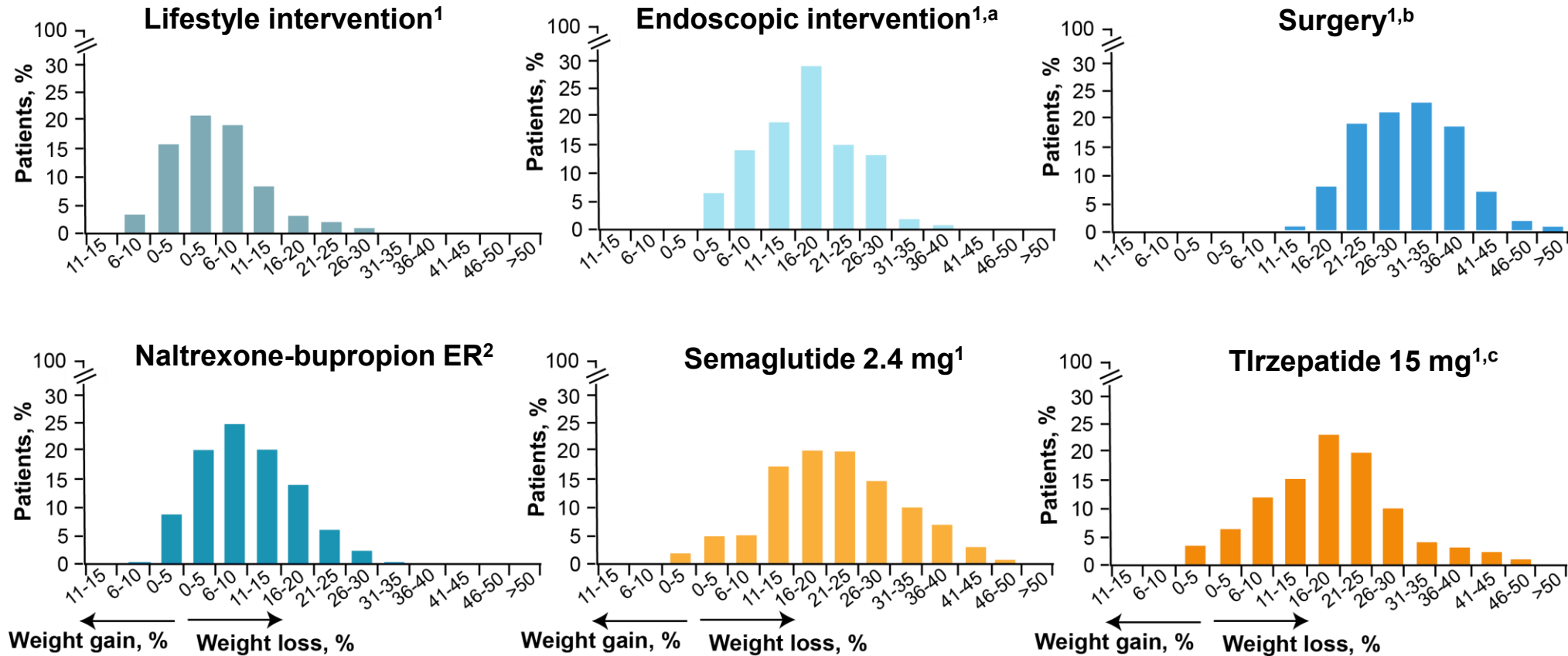


LIMITED PATIENT ACCESS³



Heterogeneity of response to weight loss interventions




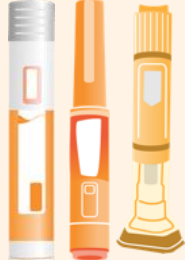
Heterogeneity in weight loss response after approximately 12 months^{1,2}



Only treatments for which 1-year data were available have been included. To enable comparison between different treatment modalities, percentage changes for total bodyweight are presented, as placebo-subtracted data are not usually reported in studies of lifestyle interventions and bariatric surgery. ^aFor endoscopic intervention, mean results for intragastric balloon and endoscopic gastroplasty are shown for which data were available. ^bFor surgery, mean results for sleeve gastrectomy and Roux-en-Y gastric bypass have been given as they account for 95% of all bariatric surgeries worldwide. ^cFor tirzepatide, since published data reached only 25% weight loss or more, the proportion of participants obtaining higher weight losses has been presumptively assigned following the unimodal distribution observed for other anti-obesity medications.

1. Perdomo CM et al. *Lancet*. 2023;401:1116-1130. 2. Data on file. Currax Pharmaceuticals LLC.

Overview of AOMs approved for long-term use

Orlistat ¹	Phentermine-topiramate ER ^{2,a}	Naltrexone-bupropion ER ³	Incretin mimetics ⁴⁻⁷
 <p>120 mg TID with meals (Rx) 60 mg TID with meals (OTC)</p>	 <p>7.5 mg/46 mg once daily 15 mg/92 mg once daily</p>	 <p>16/180 mg BID</p>	 <p>Liraglutide 3 mg QD Semaglutide 2.4 mg weekly Tirzepatide 5-15 mg weekly</p>
<p>Most common AEs^b</p> <ul style="list-style-type: none"> • Oily spotting • Flatus with discharge • Fecal urgency or incontinence • Fatty/oily stool • Oily evacuation <p>Other considerations</p> <ul style="list-style-type: none"> • Drug interactions • Decreased vitamin absorption • Oxalate nephrolithiasis 	<p>Most common AEs^b</p> <ul style="list-style-type: none"> • Paresthesia • Dizziness • Insomnia • Dry mouth <p>Other considerations</p> <ul style="list-style-type: none"> • Controlled substance • Known teratogenesis 	<p>Most common AEs^b</p> <ul style="list-style-type: none"> • GI AEs • Headache • Dizziness • Dry mouth • Insomnia <p>Other considerations</p> <ul style="list-style-type: none"> • Contraindicated in seizure disorders and uncontrolled hypertension 	<p>Most common AEs^b</p> <ul style="list-style-type: none"> • GI AEs • Headache • Fatigue • Injection site reactions <p>Other considerations</p> <ul style="list-style-type: none"> • Risk for pancreatitis • Risk for hypoglycemia • Risk for greater LBM loss?

^aOther sympathomimetic amines approved for the short-term management of obesity in the US include phendimetrazine, benzphetamine, and diethylpropion.

^bMost frequent AEs as listed in the respective product prescribing information.

AE, adverse event; AOM, antiobesity medication; GI, gastrointestinal; LBM, lean body mass.

1. Orlistat. Prescribing information. H2-Pharma LLC; 2022. 2. Qsymia. Prescribing information. Vivus LLC; 2024. 3. CONTRAVE® [prescribing information]. Nalpropion Pharmaceuticals LLC.; 2024. 4. Saxenda® Prescribing information. Novo Nordisk; 2024. 5. Wegovy® Prescribing information. Novo Nordisk; 2024. 6. Sargeant JA et al. *Endocrinol Metab.* 2019;34:247-262. 7. Zepbound™ Prescribing information. Eli Lilly and Company; 2024.

Considerations for AOM selection



Orlistat^{1,2}

Ideal use case

Patient who is not worried about GI adverse effects or is adhering to a very low fat diet^a

Special benefits

- OTC formulation available
- Lower OOP costs for patients without coverage



ER phentermine/ topiramate^{1,2}

Ideal use case

Patient with no CVD history, with history of migraine headache and no uncontrolled hypertension or risk of becoming pregnant

Special benefits

- Migraine prophylaxis
- Lower OOP costs for patients without coverage



Naltrexone- bupropion ER^{1,2}

Ideal use case

Patient with depression, tobacco use disorder, alcohol use disorder^b, and no uncontrolled hypertension

Special benefits

- Depression
- Tobacco cessation
- Lower OOP costs for patients without coverage



Incretin mimetics^{1,2}

Ideal use case

Patient with >15% weight loss clinically indicated, with CV complications, prior MI, OSA, and/or diabetes

Special benefits

- T2D
- Substantial weight loss
- OSA

CV, cardiovascular; GI, gastrointestinal; OOP, out-of-pocket; OSA, obstructive sleep apnea; T2D, type 2 diabetes.

^aAlso recommended to prescribe a daily multivitamin with orlistat owing to resulting malabsorption of fat soluble vitamins.

^bER naltrexone-bupropion contraindicated in persons undergoing abrupt discontinuation of alcohol. Monitor for mood or behavioral changes.

1. Lewis KH et al. *BMJ*. 2024;384:e072686. 2. Acosta A et al. *Obesity*. 2021;29:662-671.

Should the high prevalence of addiction disorders after bariatric surgery influence AOM selection?



AUD, alcohol use disorder; BED, binge eating disorder; LAGB, laparoscopic adjustable gastric banding; RYGB, Roux-en-Y gastric bypass.

1. Reas DL et al. *Eur Eating Disord Rev.* 2025;33:544-550. 2. Nasser K et al. *Surg. Endosc.* 2023;37:703-714. 3. Scheen AJ et al. *Diabetes Metab.* 2025;51:101612. 4. Kenkre JS et al. *Curr Obes Rep.* 2024;13:596-616.

Heterogeneity and complexity of obesity

Obesity-related Diseases

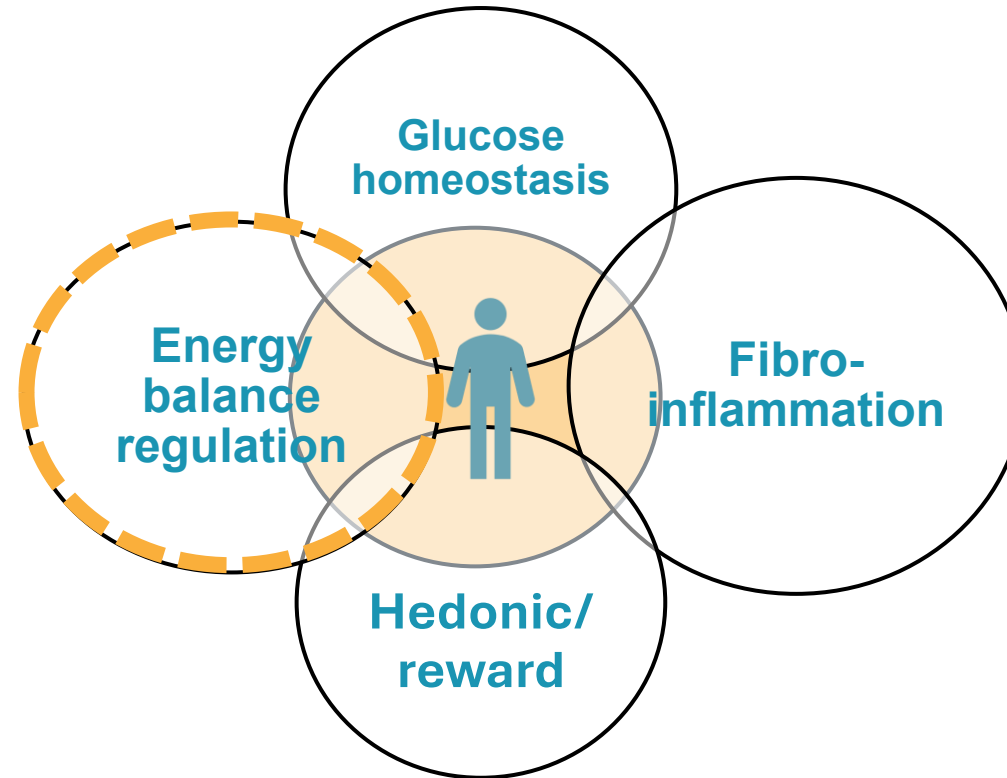
Diabetes

HTN

CVD

Cancer (some)

+280 diseases



OMICS

Exposomics

Proteomics

Metabolomics

Epi-Genetics

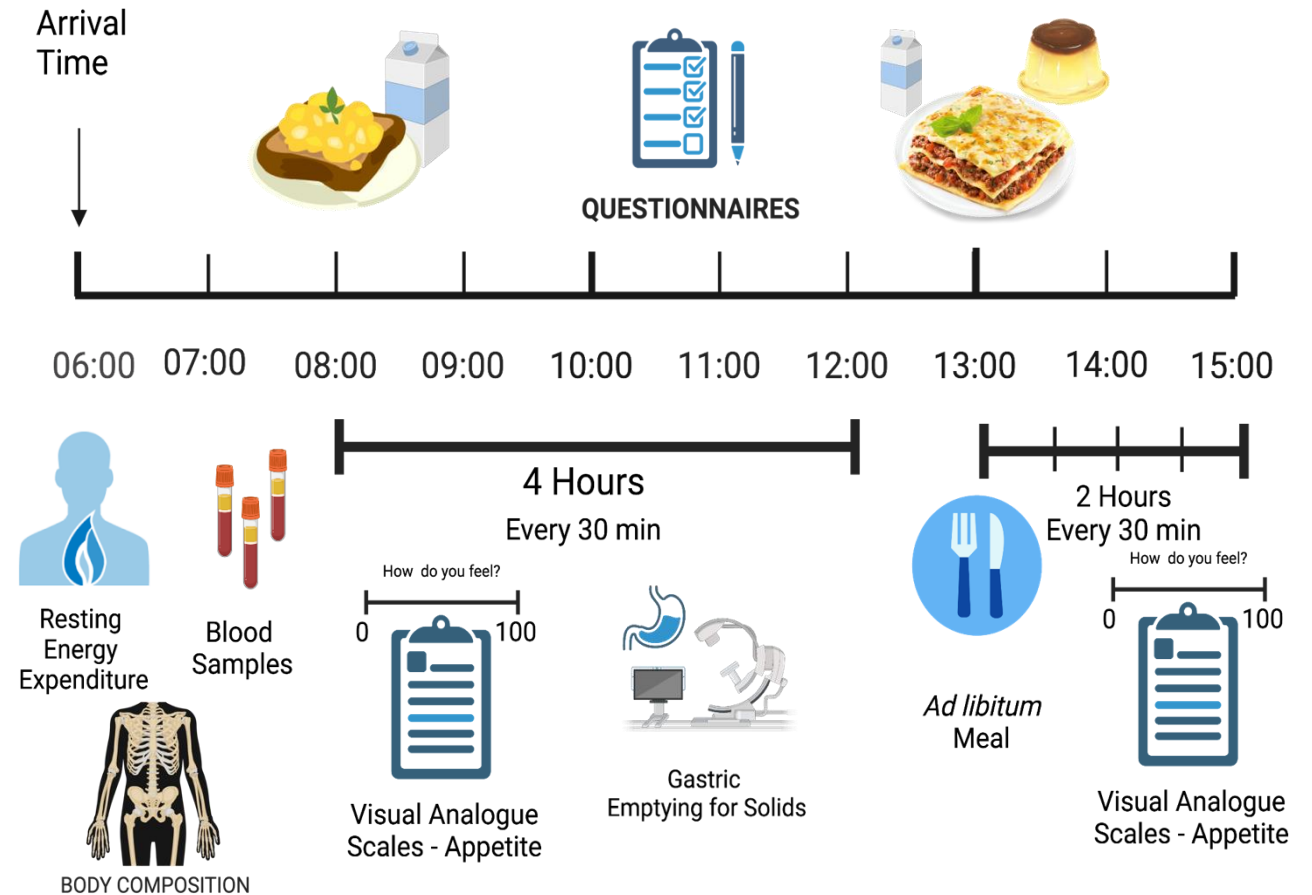
Genetics

Obesity Severity = BMI - fat mass

Social Determinants of Health

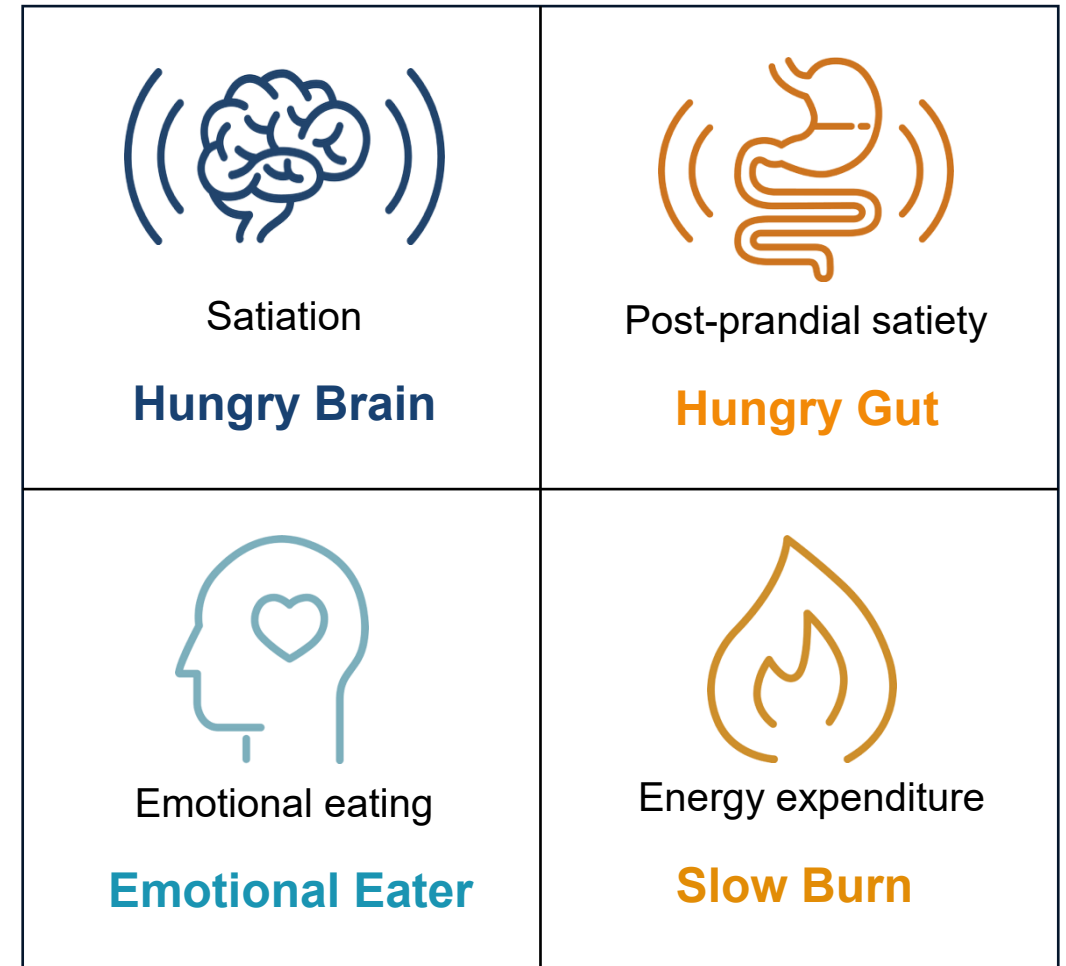
Understanding the heterogeneity of obesity through energy balance phenotypes

- We stratify obesity based on energy balance and behavioral phenotypic traits¹







Understanding the heterogeneity of obesity through energy balance phenotypes

- We stratify obesity based on energy balance and behavioral phenotypic traits¹



Understanding the heterogeneity of obesity through energy balance phenotypes

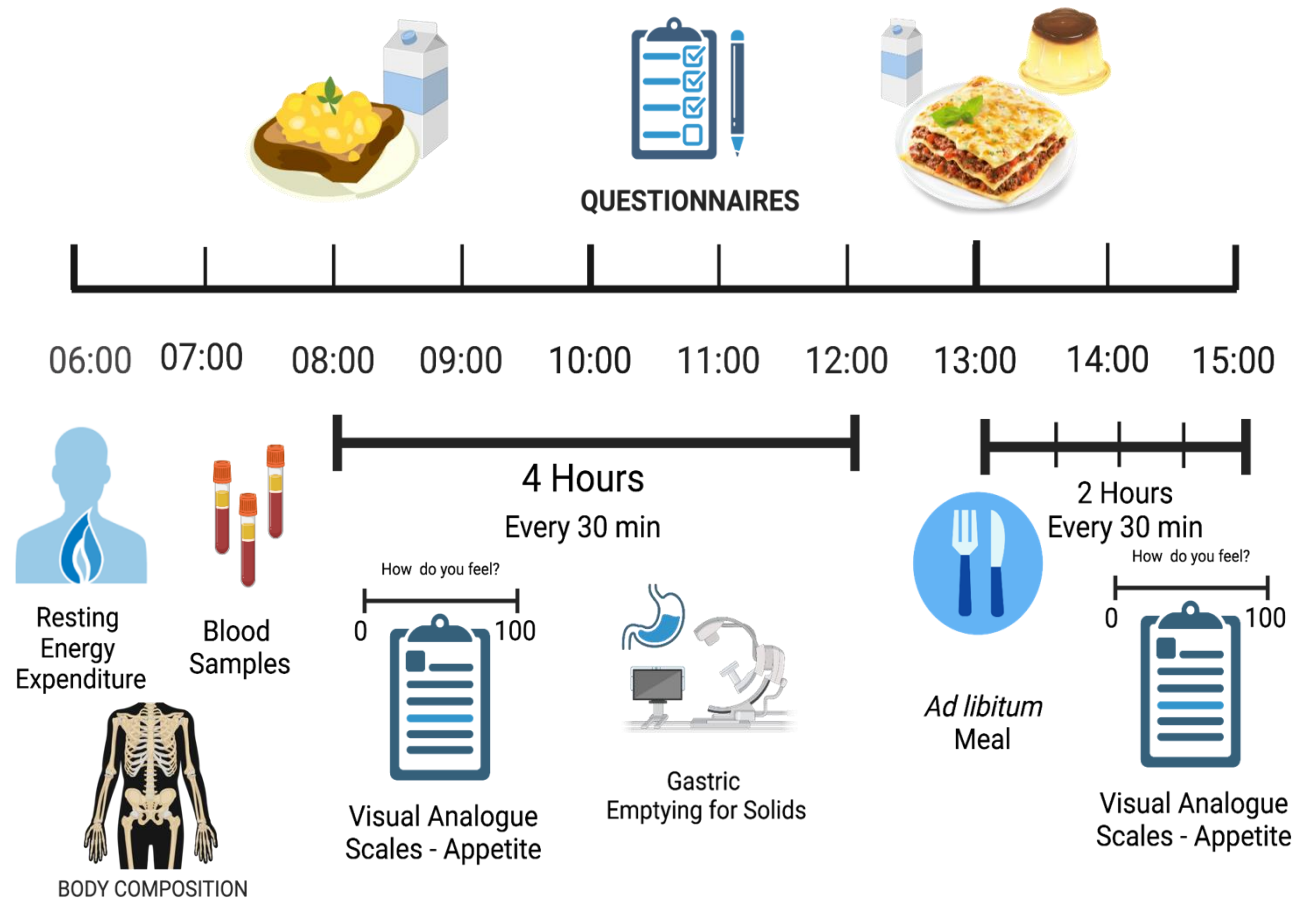
- We stratify obesity based on energy balance and behavioral phenotypic traits¹
- Antiobesity interventions that target a specific obesity phenotype improves weight loss outcomes by **1.7 to 2 fold**²⁻¹¹

	 Hungry brain	 Hungry gut	 Emotional hunger	 Slow Burn
LIFESTYLE INTERVENTION	Hungry brain diet	Hungry gut diet	Behavioral therapy Hungry feelings diet	<ul style="list-style-type: none"> • Intense exercise plan • Slow burn diet
MEDICATION ^a	Phentermine-topiramate ER	Liraglutide Semaglutide Tirzepatide	Naltrexone-bupropion ER	
ENDOSCOPY ^a	Vagal nerve block	Intragastric balloons Endoscopy sleeve gastroplasty		
SURGERY	Laparoscopic sleeve gastrectomy	Roux-en-Y gastric bypass		

1. Acosta A et al. *Gastroenterology*. 2015;148:537-546. 2. Acosta A et al. *Physiol Rep*. 2015;3(11):e12610. 3. Halawi H et al. *Lancet Gastroenterol Hepatol*. 2017;2(12):890-899. 4. Gómez V et al. *Obesity (Silver Spring)*. 2016;24(9):1849-1853. 5. Abu Dayyeh BK et al. *Clin Gastroenterol Hepatol*. 2017;15(1):37-43.e1. 6. Vargas EJ et al. *BMJ Open Gastroenterol*. 2019;6(1):e000273. 7. Acosta A et al. *Obesity (Silver Spring)*. 2021;29(4):662-671. 8. Campos A et al. *Obesity Surgery*. 2022;32(8):2632-2640. 9. Cifuentes L et al. *EClinical Medicine*. 2023;58:101923. 10. Vargas EJ et al. *Gut*. 2023;72(6):1073-1080. 11. Gala K et al. *Obesity Surg*. 2023;33(4):1284-1288.

Understanding the heterogeneity of obesity through energy balance phenotypes

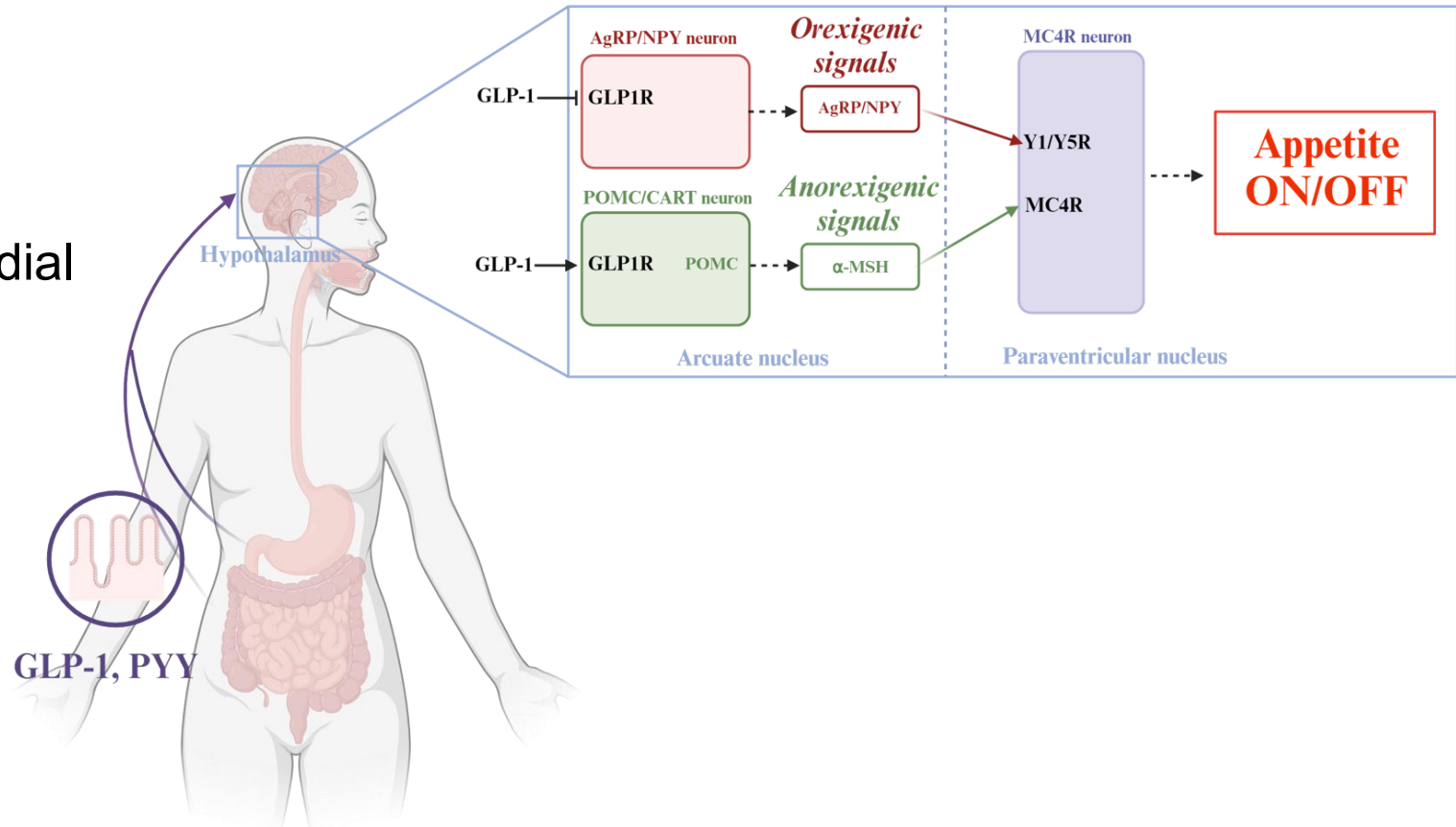
- We stratify obesity based on energy balance and behavioral phenotypic traits¹
- Antiobesity interventions that target a specific obesity phenotype improves weight loss outcomes by **1.7 to 2 fold**²⁻¹¹
- Phenotyping is not scalable; it is expensive, time-consuming and limited to a few academic centers¹²



1. Acosta A et al. *Gastroenterology*. 2015;148:537-546. 2. Acosta A et al. *Physiol Rep*. 2015;3(11):e12610. 3. Halawi H et al. *Lancet Gastroenterol Hepatol*. 2017;2(12):890-899. 4. Gómez V et al. *Obesity (Silver Spring)*. 2016;24(9):1849-1853. 5. Abu Dayyeh BK et al. *Clin Gastroenterol Hepatol*. 2017;15(1):37-43.e1. 6. Vargas EJ et al. *BMJ Open Gastroenterol*. 2019;6(1):e000273. 7. Acosta A et al. *Obesity (Silver Spring)*. 2021;29(4):662-671. 8. Campos A et al. *Obesity Surgery*. 2022;32(8):2632-2640. 9. Clfuentes L et al. *EClinical Medicine*. 2023;58:101923. 10. Vargas EJ et al. *Gut*. 2023;72(6):1073-1080. 11. Gala K et al. *Obesity Surg*. 2023;33(4):1284-1288. 12. Anazco D, Acosta A. *Int J Obes (Lond)*. 2025;49(3):452-463.

Biomarker for obesity phenotypes

- **Machine-learning gene risk score (ML-GRS) to predict appetite phenotypes**
 - Hungry Gut (abnormal postprandial satiety) and Hungry Brain (abnormal satiation)
 - 41 candidate genes from the gut-adipose-brain pathway

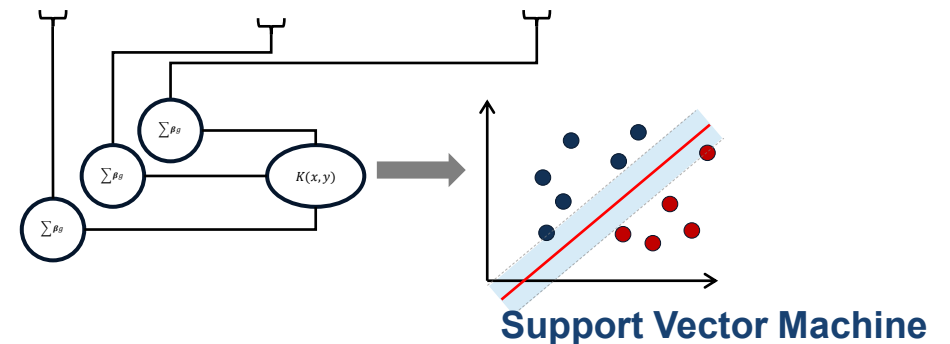
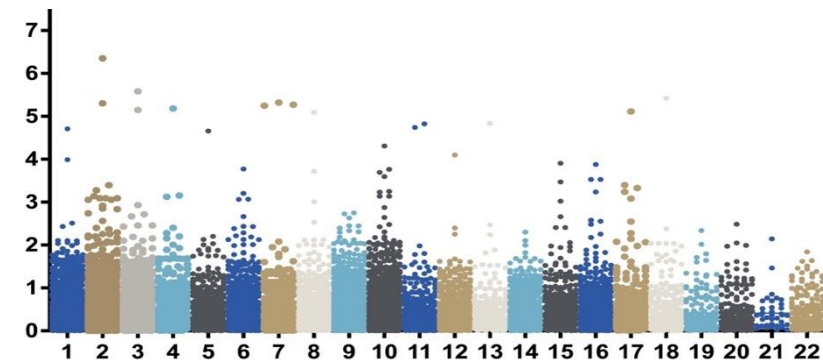


Biomarker for obesity phenotypes

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 - Hungry Gut (abnormal postprandial satiety) and Hungry Brain (abnormal satiation)
 - 41 candidate genes from the gut-adipose-brain pathway
 - Using machine-learning techniques, the most informative GRS were selected to create the biomarker to predict hungry gut phenotype

Development of ML-GRS

Manhattan Plot – GWAS

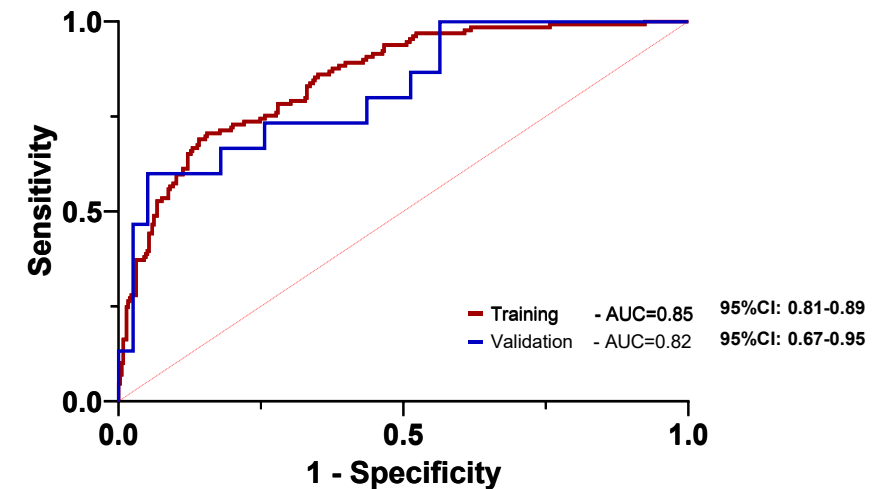


Biomarker for obesity phenotypes

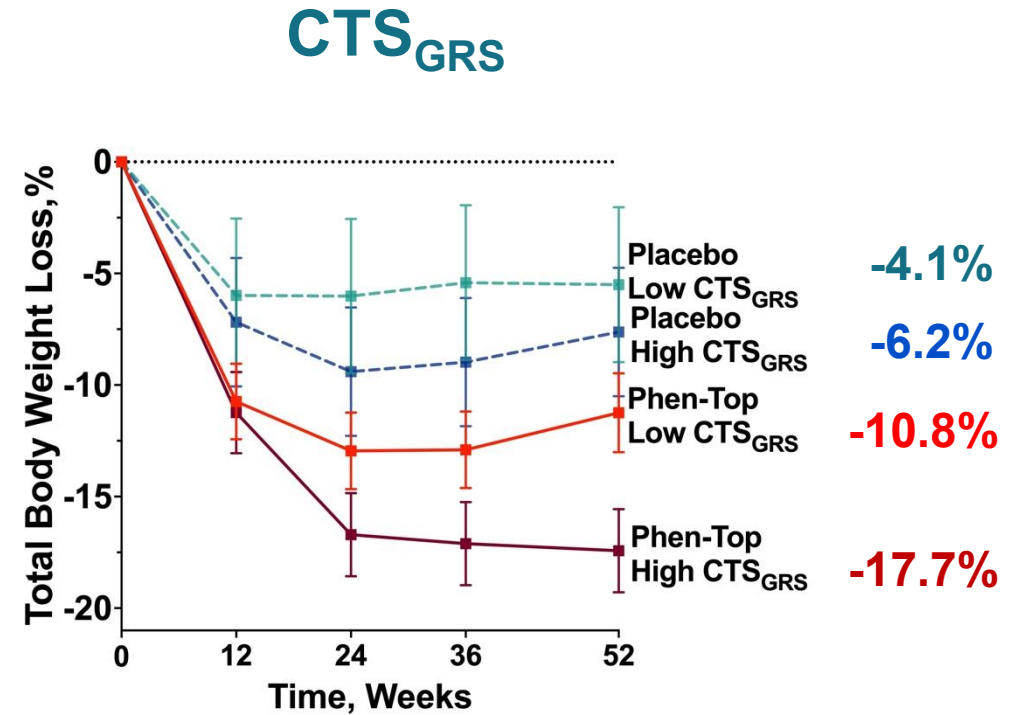
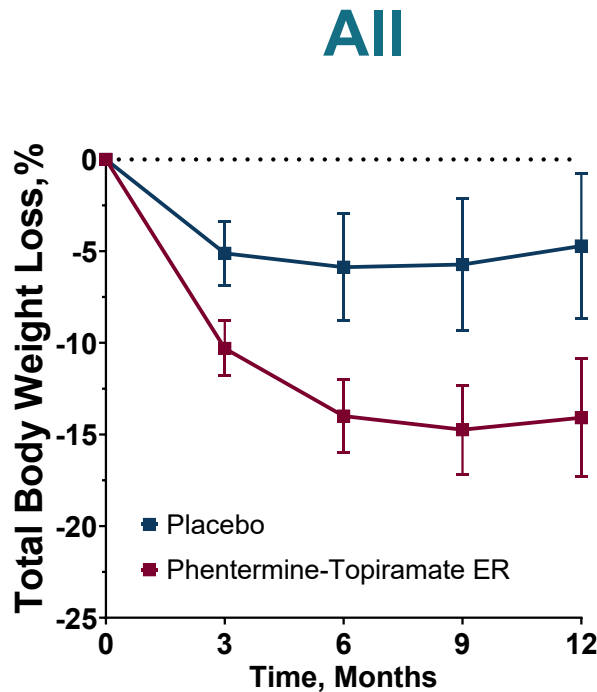
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 - Using machine-learning techniques, the most informative GRS were selected to create the biomarker to predict hungry gut phenotype

Performance of the ML-GRS model to predict appetite phenotype

- Hungry Gut Positive (HG+), as CTS - ML-GRS < 0.50
- Hungry Brain Positive (HG+), as CTS - ML-GRS > 0.50



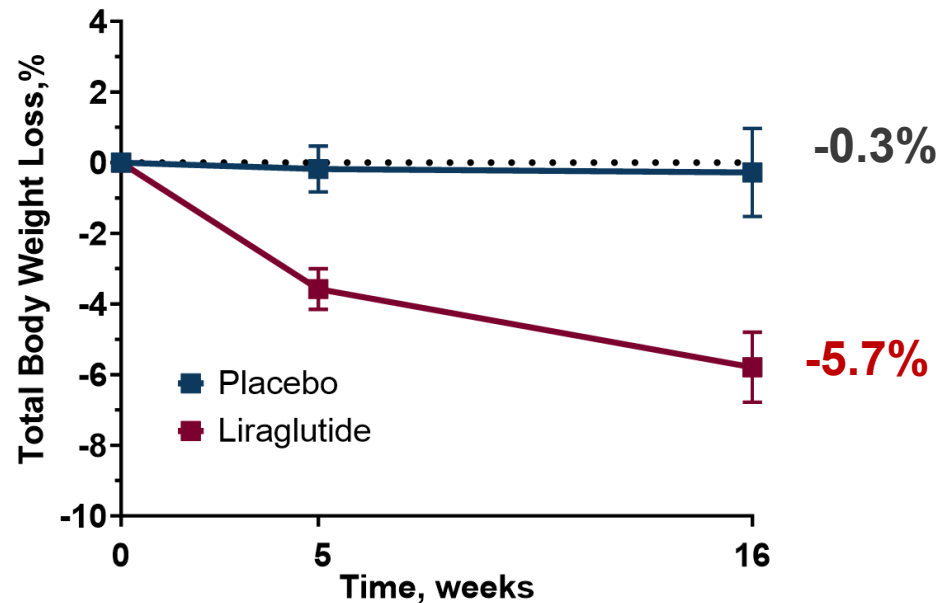
Satiation or CTS_{GRS} prediction of response to phentermine-topiramate ER



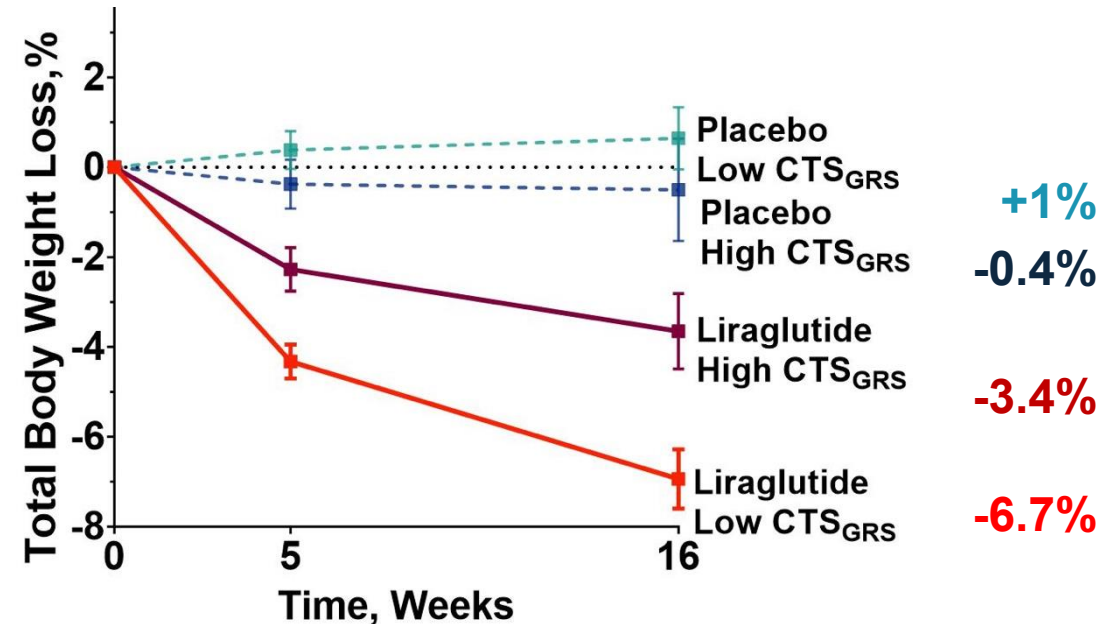
80 patients (BMI: 38 ± 6.9 kg/m²)
 From placebo-controlled, 52-week trial of
 Phentermine-topiramate ER 7.5/46mg daily

CTS_{GRS} prediction of response to liraglutide

All participants



CTS_{GRS}



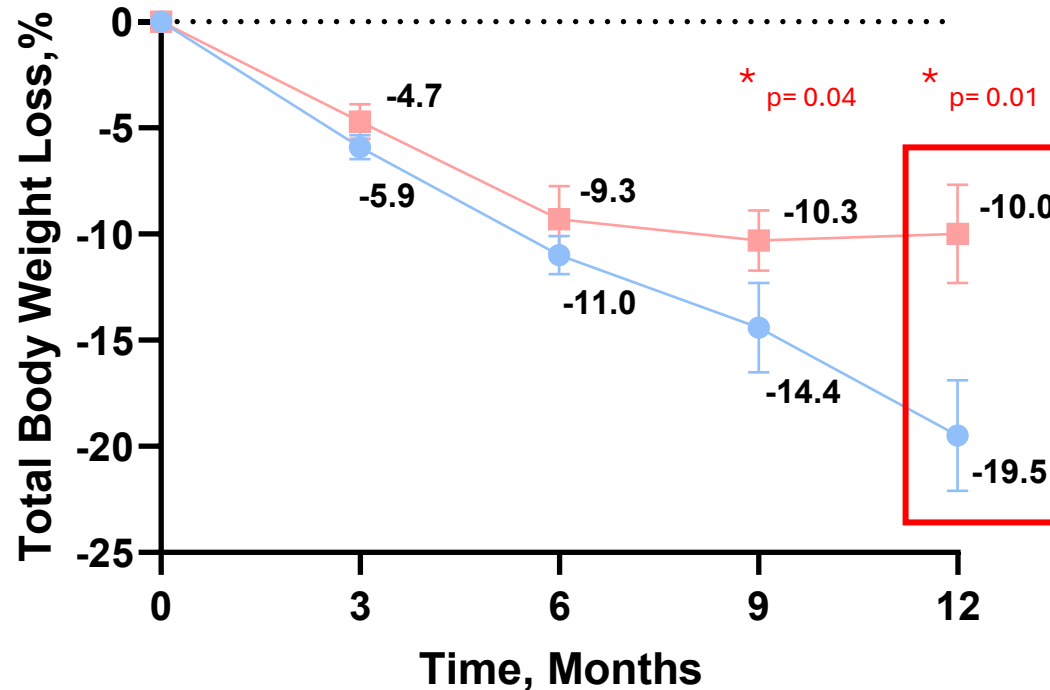
Randomized placebo-controlled, double-blinded trial¹
Liraglutide 3 mg SC daily for 16 weeks
N= 121 patients (BMI: 39.09 ± 7.40 kg/m²)

CTS, calories to satisfaction; GRS, genetic risk score; SC, subcutaneous.

1. Maselli et al. *Obesity*. 2022;30(8):1608-1620. 2. Cifuentes L et al. *Cell Metab*. 2025;37(8):1655-1666.e5.

Clinical utility: hungry gut test and weight loss outcomes to semaglutide

- Prospective cohort real-world evidence of semaglutide 2.4 mg SC weekly
- N=84 patients (BMI: 38 ± 7.40 kg/m²)



■ CTS_{GRS} >0.50 (high)
Hungry gut-negative

■ CTS_{GRS} <0.50 (low)
Hungry gut-positive

2-fold higher weight loss associated with hungry gut-positive at 12 months compared with hungry gut-negative

	n/N ^a				
	Start	3 months	6 months	9 months	12 months
Hungry Gut +	51/51	36/50	40/45	24/32	19/22
Hungry Gut -	33/33	26/33	24/30	24/27	16/24

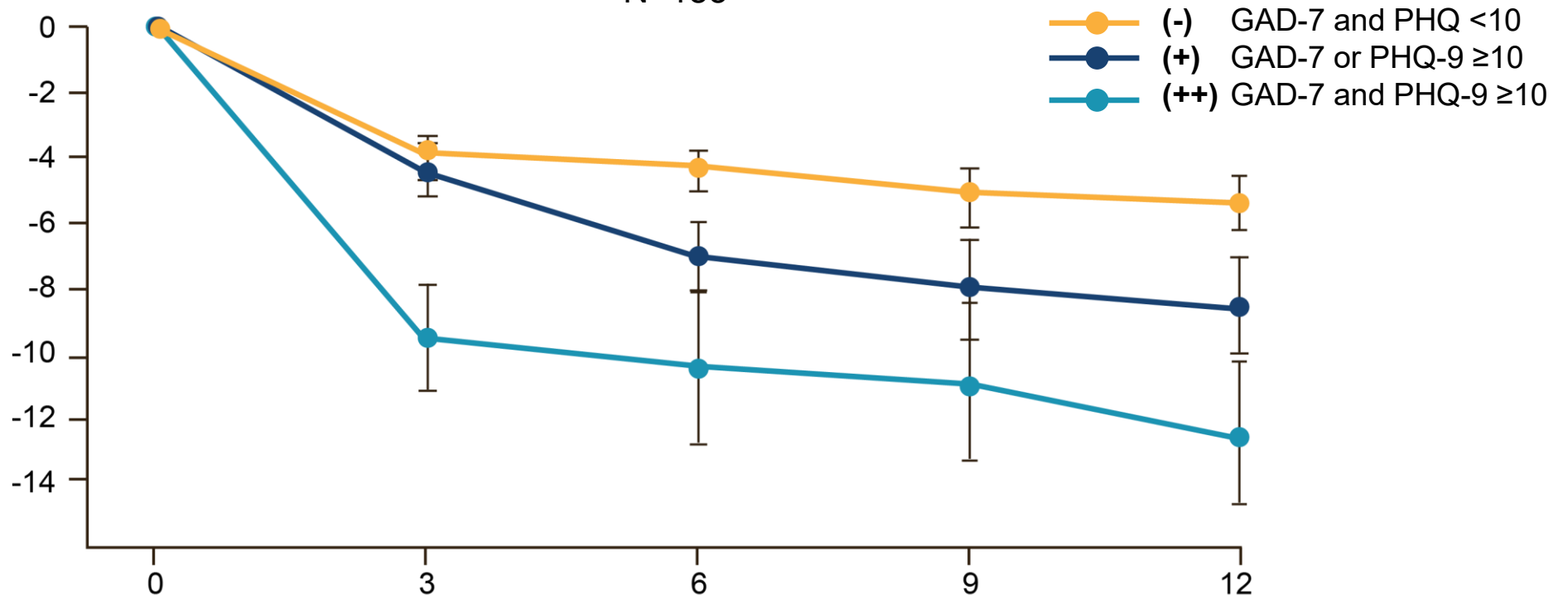
CTS, calories to satisfaction; GRS, genetic risk score; SC, subcutaneous.
^aNumber of participants with weight values/number of possible participants.
 Fansa S et al. Presented at ADA. 2024.

Emotional hunger phenotype: questionnaires for clinical validity of naltrexone-bupropion ER

12-month weight loss outcomes in patients with high vs low depression/anxiety levels treated with naltrexone-bupropion ER

Retrospective, 12-month, real-world evidence

N=185



185 patients
Mean age 52.1 ±13.9 y
83% female
BMI 39.4 ± 7.5 kg/m²

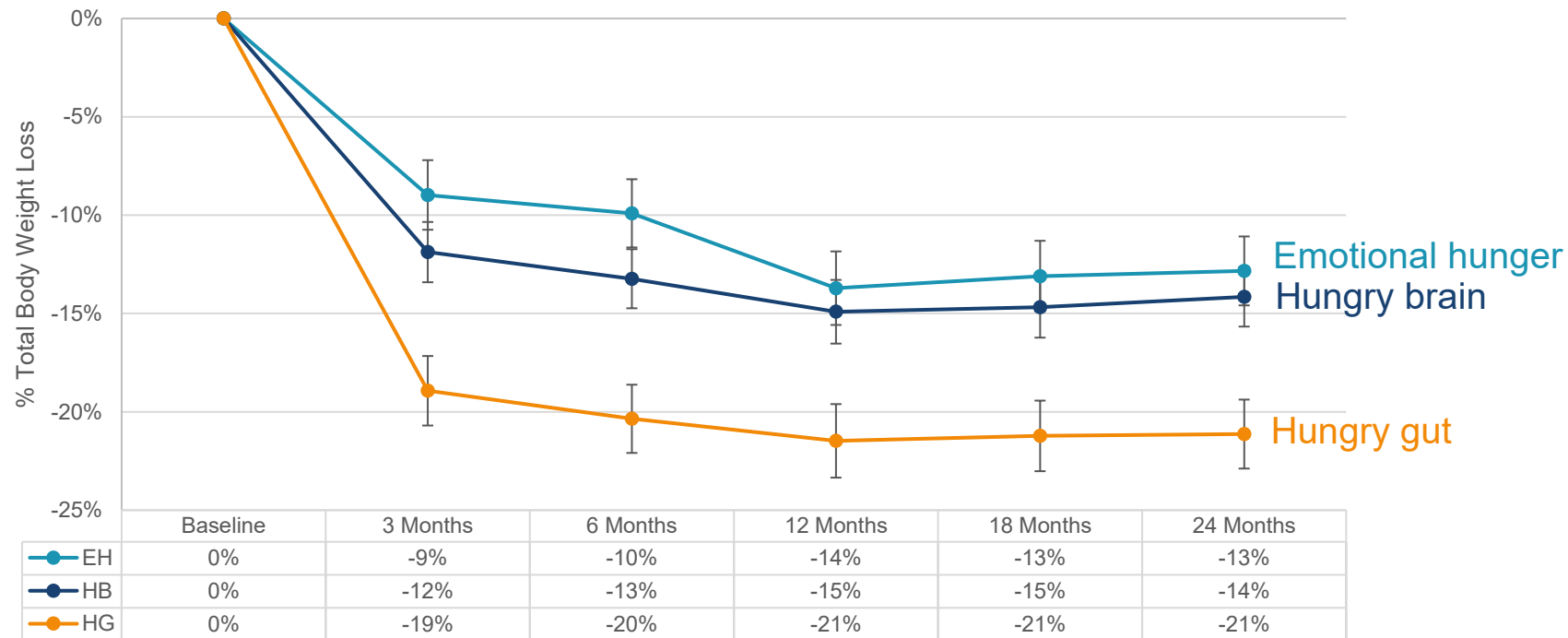
Obesity phenotypes and endoscopic sleeve gastroplasty: multicenter study

Participant demographics (n=49)

	Emotional hunger (n=14)	Hungry brain (n=19)	Hungry gut (n=16)
Female (n, %)	12 (85.7%)	15 (84.2%)	15 (93.8%)
Age at Procedure, years	50.5 (3.1)	48.1 (2.6)	44.5 (2.9)
BMI at Procedure, kg/m ²	38.6 (5.1)	37.3 (4.6)	38.3 (4.2)
Comorbidities			
Type 2 diabetes mellitus	3 (21.4%)	3 (15.8%)	0 (0.0%)
Anxiety	5 (35.7%)	7 (36.8%)	5 (31.3%)
Depression	5 (35.7%)	4 (21.1%)	8 (50.0%)
Hypertension	4 (28.6%)	4 (21.1%)	1 (6.3%)
Hyperlipidemia	4 (28.6%)	6 (31.6%)	6 (37.5%)
OSA	2 (14.3%)	5 (26.3%)	4 (25.0%)

Obesity phenotypes and endoscopic sleeve gastroplasty

Endoscopic sleeve gastroplasty



Hungry Gut Positive (Low $CTS_{GRS} < 0.50$) are best responders to ESG.

RYGB & genetics: leptin-melanocortin pathway

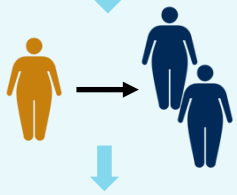
Methods



50,000 participants of the Mayo Clinic Biobank were assessed for patients with history of RYGB



Eligible patients were genotyped for heterozygous variants in the leptin-melanocortin pathway

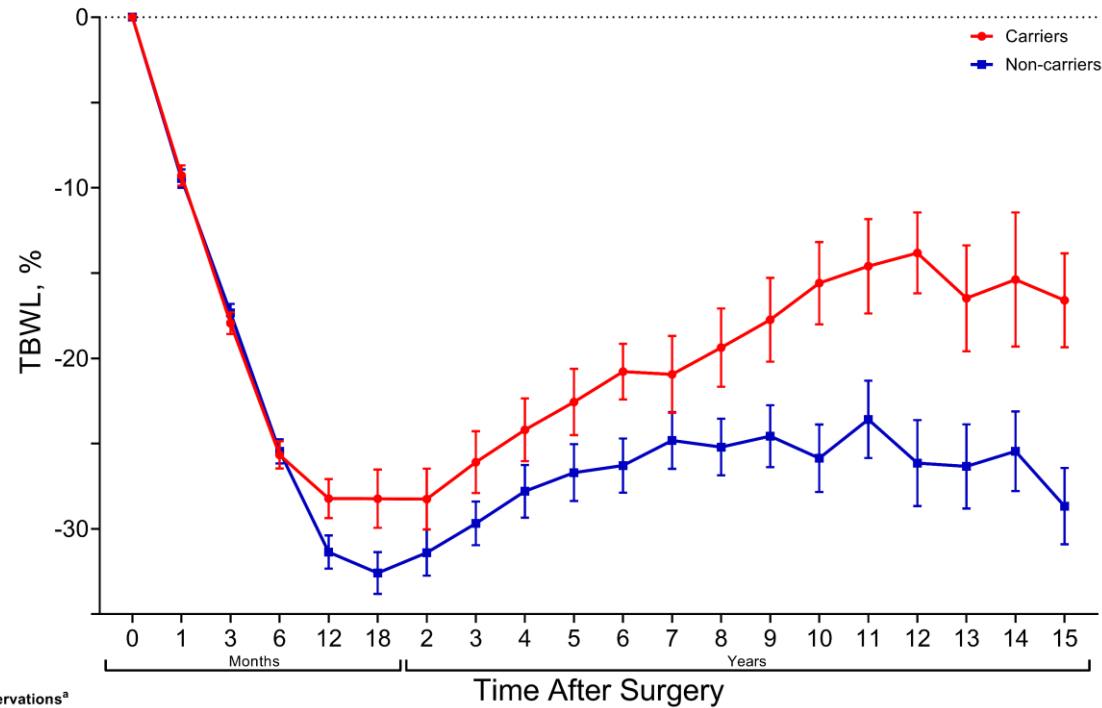


Each carrier of a heterozygous variant was randomly matched with 2 non-carrier controls based on: sex, age, BMI, & years since surgery.

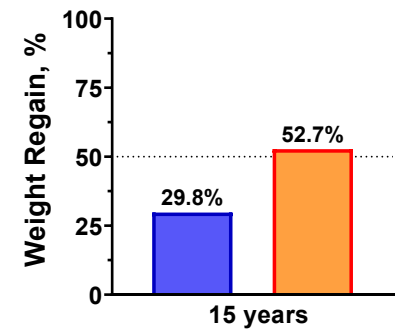
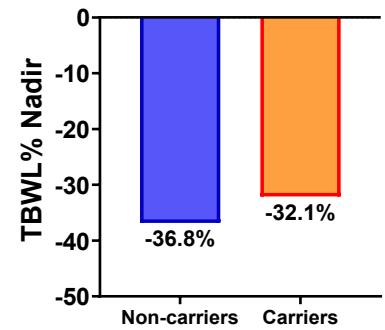


Data was abstracted from the electronic medical record for up to 15 years after surgery

RYGB and carriers of leptin-melanocortin mutations



	0	1	3	6	12	18	2	3	4	5	6	7	8	9	10	11	12	13	14	15
Observations ^a	50	43	44	46	48	37	39	39	36	34	37	25	25	21	23	18	18	15	14	15
Possible Observations ^b	50	50	50	50	50	49	48	46	46	45	40	35	32	27	26	23	22	19	18	16
Carriers																				
Non-Carriers	100	78	88	94	93	80	83	86	82	73	68	66	65	59	50	49	40	39	36	33
	100	100	100	100	100	99	99	97	94	90	88	86	77	72	68	59	51	50	45	43

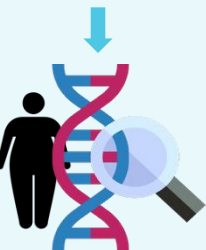


Obesity phenotypes and bariatric surgery–RYGB

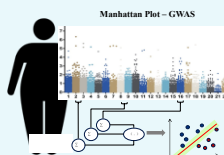
Methods



50,000 participants of the Mayo Clinic Biobank were assessed for patients with history of RYGB



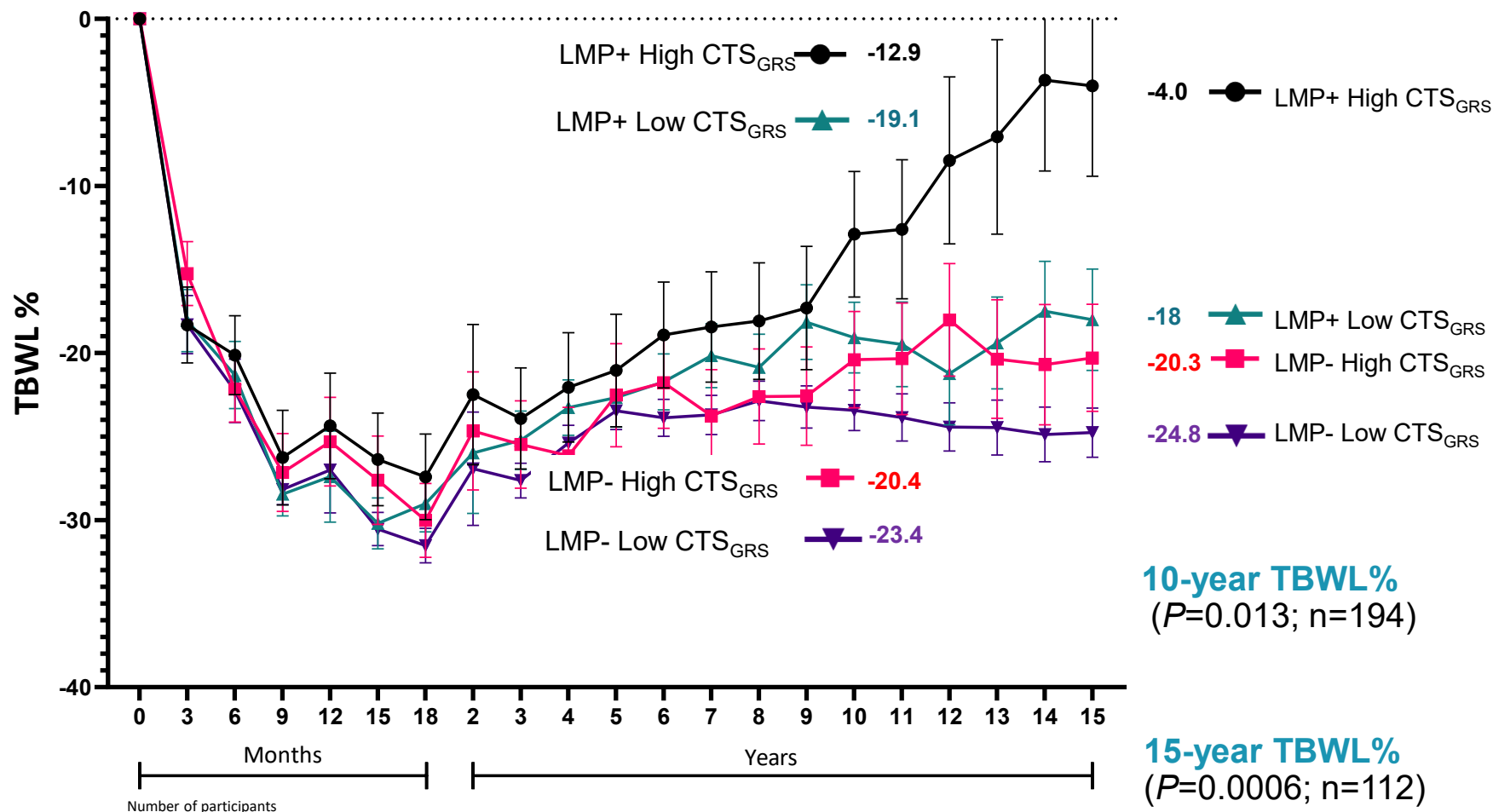
Eligible patients were genotyped for heterozygous variants in the leptin-melanocortin pathway (LMP+ or LMP-)



Calories to Satiation Gene Risk Score was calculated (CTS_{GRS} low or high)



Data was abstracted from the electronic medical record for up to 15 years after surgery



	0	3	6	9	12	15	18	2	3	4	5	6	7	8	9	10	11	12	13	14	15	
LMP+/CTS _{GRS} ⁺	15	10	14	6	15	10	14	14	13	10	10	12	12	12	11	9	9	6	5	5	4	
LMP+/CTS _{GRS} ⁻	28	11	24	9	27	11	19	21	18	13	12	16	17	18	17	16	14	14	14	14	12	12
LMP-/CTS _{GRS} ⁺	56	37	47	30	46	34	33	42	40	40	37	44	37	38	31	29	25	16	23	17	13	
LMP-/CTS _{GRS} ⁻	254	157	185	101	193	130	148	181	170	145	143	158	154	163	149	140	117	111	98	92	84	

CTS, calories to satisfaction; LMP, leptin-melanocortin pathway; RYGB, Roux-en-Y gastric bypass; TBWL, total body weight loss.

Data are presented as LSM ± SEM.

Espinosa MA et al.... Acosta A. Presented at DDW 2025.

Sleeve & genetics: leptin-melanocortin pathway

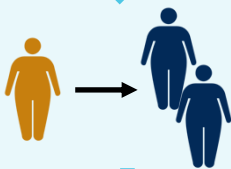
Methods



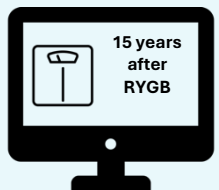
50,000 participants of the Mayo Clinic Biobank were assessed for patients with history of RYGB



Eligible patients were genotyped for heterozygous variants in the leptin-melanocortin pathway

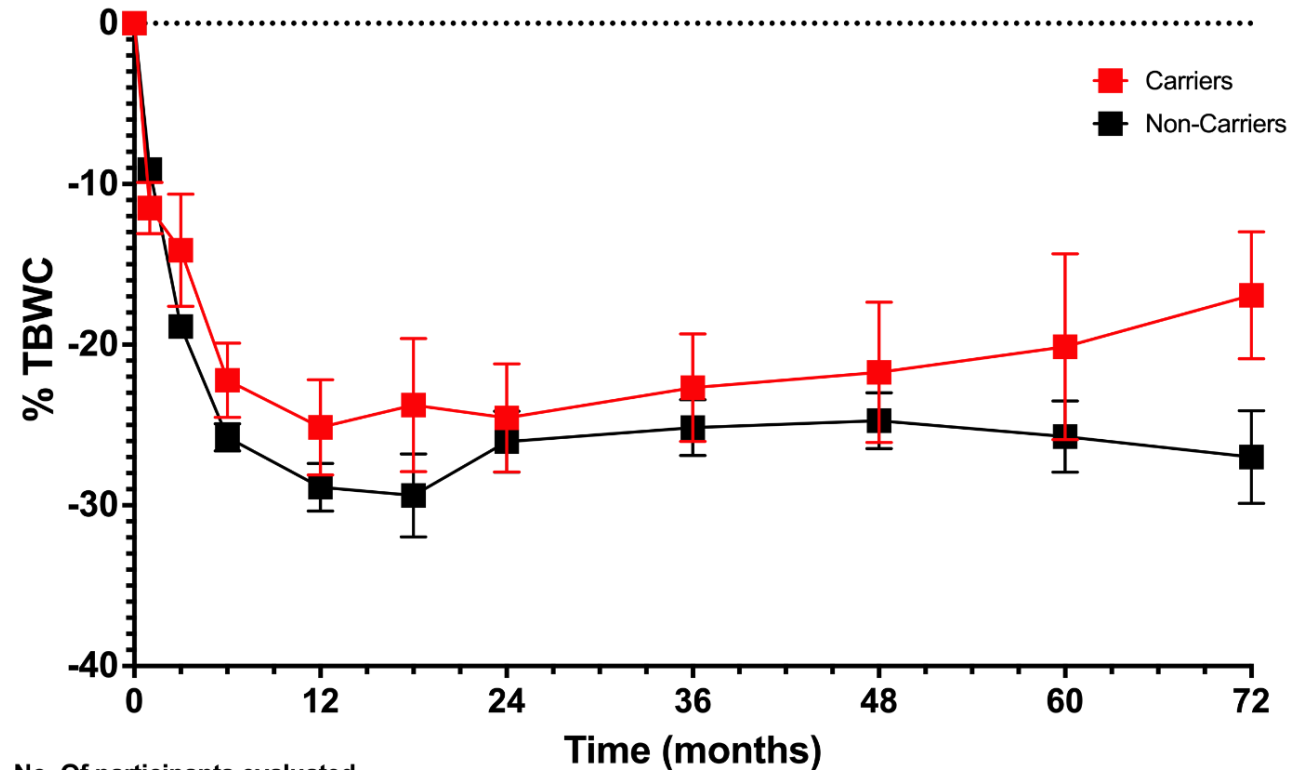


Each carrier of a heterozygous variant was randomly matched with 2 non-carrier controls based on: sex, age, BMI, & years since surgery



Data was abstracted from the electronic medical record for up to 6 years after surgery

Sleeve and carriers of leptin-melanocortin mutations



No. Of participants evaluated

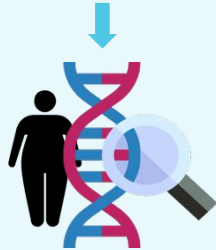
Carriers	17	18	18	15	18	18	15	12	14
Non-carriers	26	25	25	16	26	24	23	17	12

TORe- RYGB & genetics: leptin-melanocortin pathway

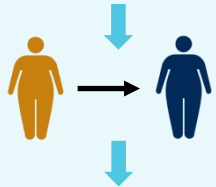
Methods



50,000 participants of the Mayo Clinic Biobank were assessed for patients with history of RYGB



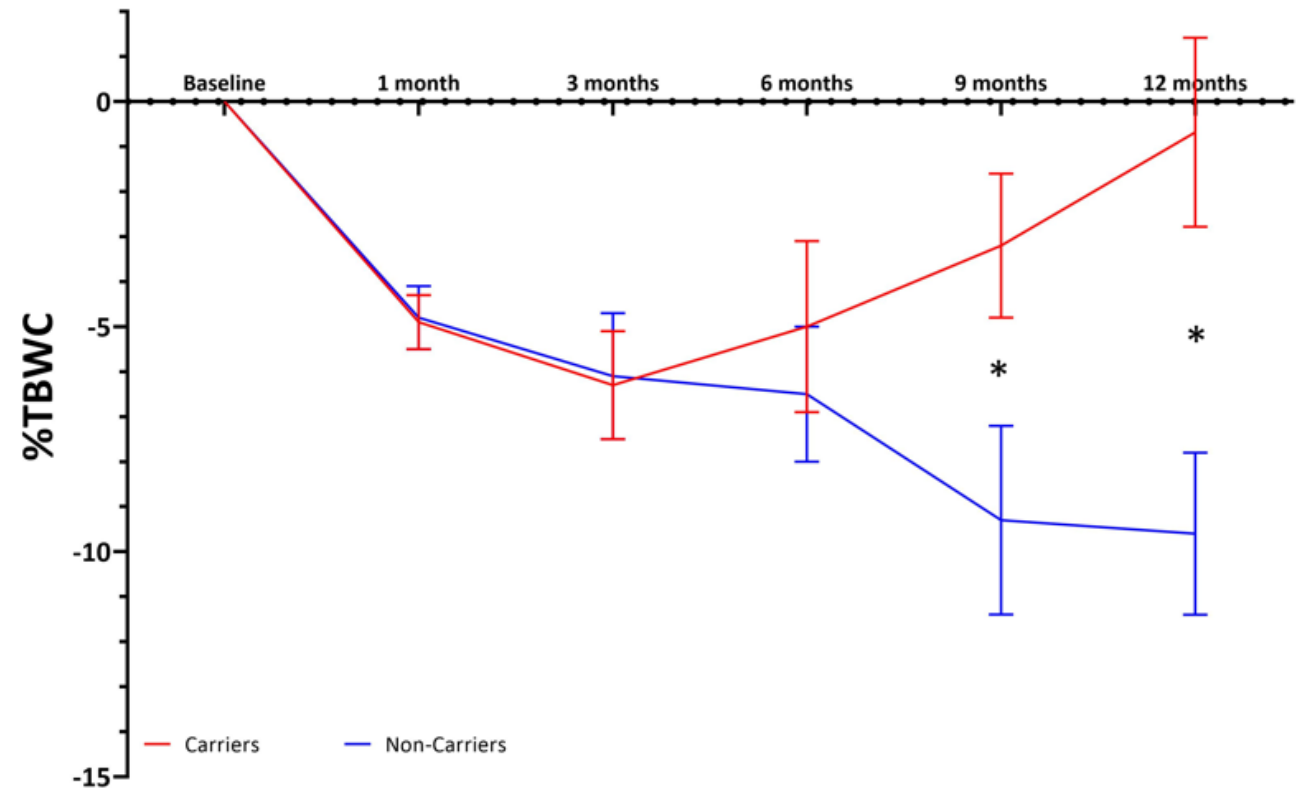
Eligible patients were patients with RYGB plus TORe who genotyped for heterozygous variants in the leptin-melanocortin pathway



All participants including. No matching.



Data was abstracted from the electronic medical record for up to 1 year after procedure



Carriers (n)	19	14	14	11	13
Non-Carriers (n)	28	23	21	12	20

Precision Obesity: cost-effective-phenotype based obesity solution



Hungry brain



Hungry gut



Emotional hunger



Slow Burn

	Hungry brain	Hungry gut	Emotional hunger	Slow Burn
LIFESTYLE INTERVENTION	Hungry brain diet	Hungry gut diet	Behavioral therapy Hungry feelings diet	<ul style="list-style-type: none"> • Intense exercise plan • Slow burn diet
MEDICATION^a	Phentermine-topiramate ER	Liraglutide Semaglutide Tirzepatide	Naltrexone-bupropion ER	
ENDOSCOPY^a	Vagal nerve block	Intragastric balloons Endoscopy sleeve gastroplasty		
SURGERY	Laparoscopic sleeve gastrectomy	Roux-en-Y gastric bypass		

^aValidated in randomized clinical trials.

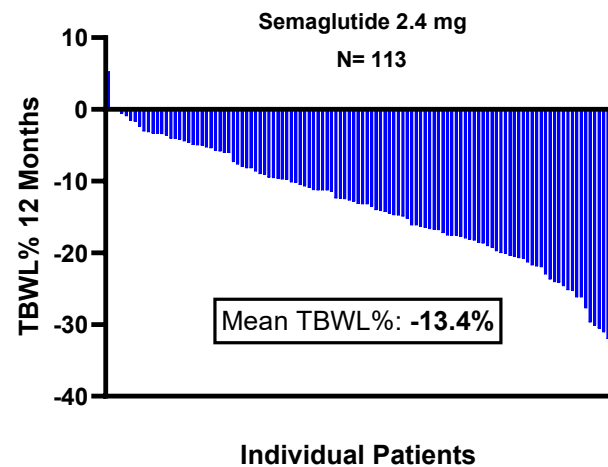
1. Acosta A et al. *Gastroenterology*. 2015;148:537-546. 2. Acosta A et al. *Physiol Rep*. 2015;3(11):e12610. 3. Halawi H et al. *Lancet Gastroenterol Hepatol*. 2017;2(12):890-899. 4. Gómez V et al. *Obesity (Silver Spring)*. 2016;24(9):1849-1853. 5. Abu Dayyeh BK et al. *Clin Gastroenterol Hepatol*. 2017;15(1):37-43.e1. 6. Vargas EJ et al. *BMJ Open Gastroenterol*. 2019;6(1):e000273. 7. Acosta A et al. *Obesity (Silver Spring)*. 2021;29(4):662-671. 8. Campos A et al. *Obesity Surgery*. 2022;32(8):2632-2640. 9. Cifuentes L et al. *EClinical Medicine*. 2023;58:101923. 10. Vargas EJ et al. *Gut*. 2023;72(6):1073-1080. 11. Gala K et al. *Obesity Surg*. 2023;33(4):1284-1288. 12. Cifuentes L et al. *Cell Metab*. 2025;37(8):1655-1666.e5.

Key take-away points

- “One-treatment-fits-all” is not working
- Obesity is a complex and heterogenous disease with multiple phenotypes
- Phenotype-guided classification and interventions double weight loss
- Biomarker predicts phenotypes and identifies best responders to obesity interventions

Opportunities to tackle ~~Obstacles~~ In the obesity pandemic

HETEROGENEOUS OUTCOMES¹



HIGH COST²



LIMITED PATIENT ACCESS³



Better outcomes with
MyPhenome Test (CTS_{GRS})

Obesity Rx cost-effective?

Increase access
and reduce stigma?

Precision Medicine for Obesity Program

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Matthew Clark



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Qualitative Health Sciences	Janet Olson Suzette Bielinski Ryan Lennon

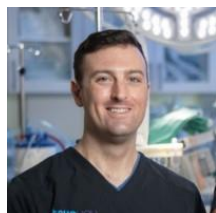
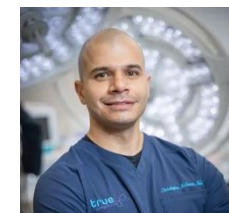
Phenomix Sciences

Tim O'Connor, PhD



True You Weight Loss

Chris McGowan Dan Maselli



Case Study and Q&A



Thank you!

