# Effectiveness and durability of common weight loss methods

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## Background

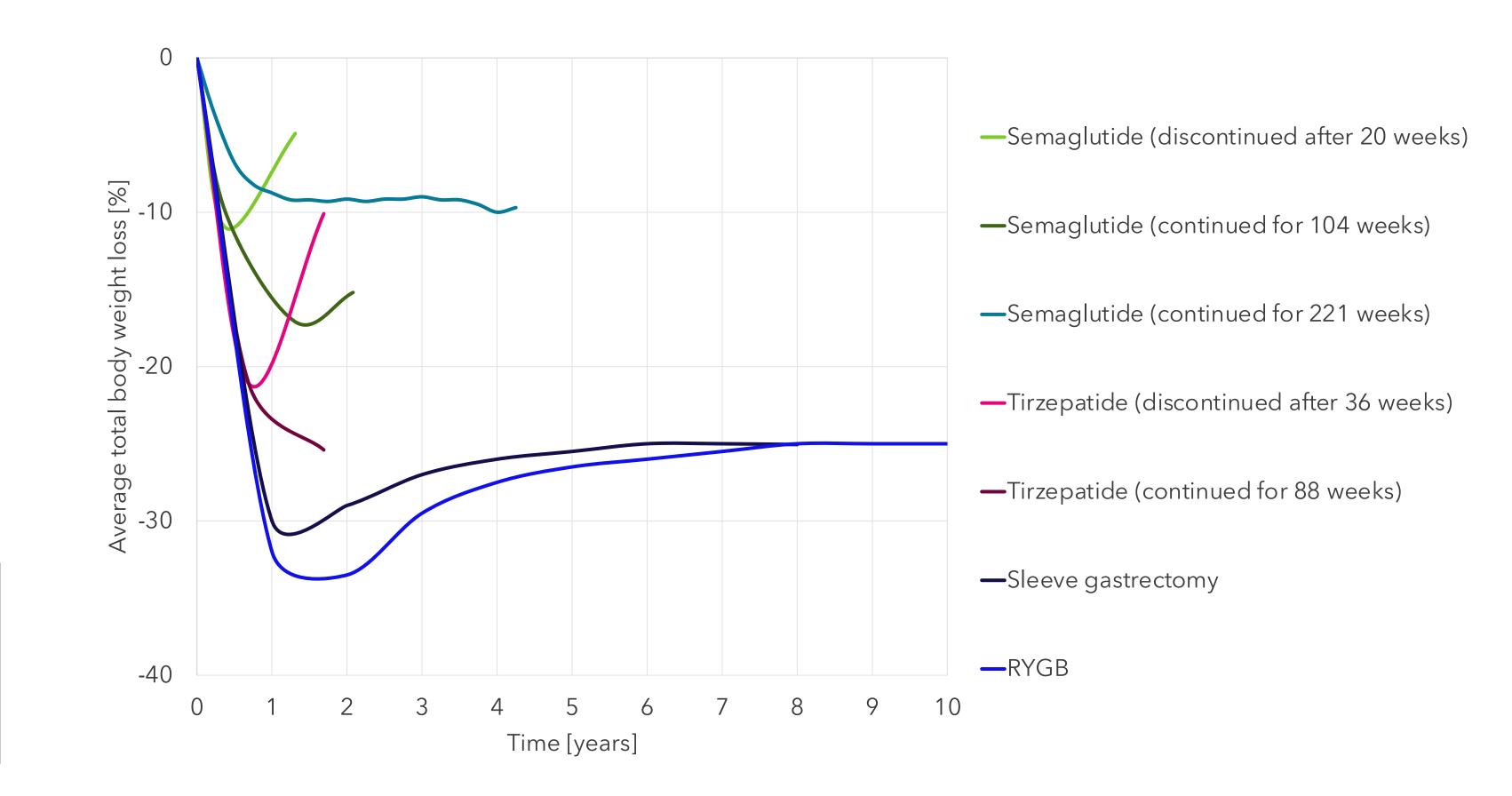
• Obesity is a chronic disease requiring long-term treatment to achieve sustainable weight loss.<sup>1</sup>

## Methods

- Based on published data, three treatment approaches were compared regarding their effectiveness and stability of results (Table 1):
  - Lifestyle interventions<sup>2</sup>
  - GLP-1 receptor agonists (semaglutide and tirzepatide)<sup>3-7,9</sup>
  - Bariatric surgery (Roux-en-Y gastric bypass (RYGB) and sleeve gastrectomy)<sup>8</sup>

### **Table 1 Publication overview**

Intervention	Reference	Study type	Population size	Nadir weight with treatment	Weight loss at plateau or end of study with treatment
Lifestyle interventions	Nordmo et al. (2020) <sup>2</sup>	Systematic review	723	7.4% after treatment of varying lengths	Regain of lost weight on average within 4.1 years
Semaglutide	Rubino et al. (2021) <sup>3</sup>	RCT (STEP-4)	803	-17.8% at 15 months	-17.1% at 17 months
Semaglutide	Garvey et al. (2022) <sup>5</sup>	RCT (STEP-5)	304	-17.5% at 17 months	-15.2% at 26 months
Semaglutide	Ryan et al. (2024) <sup>9</sup>	RCT (SELECT)	8,803	-10.0% at 4 years	-9.7% at 4.3 years
Semaglutide	Wilding et al. (2021) <sup>6</sup>	RCT (STEP-1)	1,961	-16% at 15 months	-14.9 at 17 months
Tirzepatide	Aronne et al. (2024) <sup>4</sup>	RCT (SURMOUNT-4)	670	-25.9% at 19 months	-25.8% at 22 months
Tirzepatide	Jastreboff et al. (2022) <sup>7</sup>	RCT (SURMOUNT-1)	2539 Tirzepatide: <b>5 mg</b> : 630 <b>10 mg</b> : 636 <b>15 mg</b> : 630	Tirzepatide at 18 months: 5 mg: -16.0% 10 mg:-21.4% 15 mg: -22.5%	Tirzepatide at 18 months: 5 mg: -16.0% 10 mg: -21.4% 15 mg: -22.5%
RYGB	van Rijswijk et al. (2021) <sup>8</sup>	Systematic review	18,665	-34% at 1.5 years	-25% at 8 years
Sleeve gastrectomy	van Rijswijk et al. (2021) <sup>8</sup>	Systematic review	6,095	-31% at 1.3 years	-25% at 6 years



## Figure 1 Percentage of total body weight lost over time combined from different publications.

The semaglutide graphs are based on the STEP trials and the SELECT trial where participants received a weekly dose of 2.4 mg semaglutide for 20 weeks and 221 weeks, respectively.<sup>3,9</sup> This treatment was then either continued through week 68 (STEP-1, STEP-4)<sup>3,6</sup> to week 104 (Step-5),<sup>5</sup> to week 221 (SELECT),<sup>9</sup> or discontinued and switched to placebo until week 68 (STEP-4).<sup>3</sup> The tirzepatide graphs are based on the SURMOUNT-4 trial, where participants were treated with tirzepatide for 36 weeks.<sup>4</sup> This treatment was then either continued or discontinued and switched to placebo until week 88.<sup>4</sup> Finally, the data for sleeve gastrectomy and Roux-en-Y gastric bypass (RYGB) is based on the pooled mean percentage of total weight loss from 46 studies in a systematic review.<sup>8</sup> The longest follow-up times were 8 years for sleeve gastrectomy and 10 years for RYGB.<sup>8</sup>

### Disclaimer

MEJ, CF, GP, and MK declare no conflict of interest.

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## Conclusion

- Lifestyle interventions do not lead to sustainable weight loss.<sup>2</sup>
- The effect of weight loss medications is durable only with sustained use of the medications<sup>3,4,9</sup>
- Bariatric surgery demonstrated the greatest weight loss and sustained weight loss in comparison to other interventions.<sup>8</sup>

### Results

- A systematic review of lifestyle interventions demonstrated a 7.4% mean weight loss at the end of the treatment period, but 0.14% of weight was regained per month post-treatment, reaching pre-intervention weights within 4.1 years.<sup>2</sup>
- Weekly injections of semaglutide for 20 weeks and tirzepatide for 36 weeks resulted in 10.6% and 21.1% weight loss, respectively.<sup>3,4</sup>
- Once injections were stopped, approximately half of that weight was regained within a year. $^{3,4}$
- If injections were continued, a plateau was reached after 17-18 months at 22.5% for tirzepatide<sup>7</sup> and between 9.7% and 14.9% for semaglutide.<sup>6,9</sup>
- Within 2 years of semaglutide treatment, approximately 2% of the lost weight was regained in one study.<sup>5</sup> In another study, weight loss plateaued at approx. 9.5% over 4 years.<sup>9</sup>
- A systematic review of weight loss outcomes after RYGB and sleeve gastrectomy demonstrated a total weight loss of 31.9% and 29.5% one year after surgery, respectively.8
- Over the following years, roughly 5-7% of that weight was regained, so that both procedures resulted in a stable total weight loss of approximately 25% after 8-10 years.<sup>8</sup>

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