

# Impact of Digital Engagement on Weight Loss Outcomes in Obesity Management: A Retrospective Service Evaluation of GLP-1 and Dual GLP-1/GIP Receptor Agonist Therapy in the United Kingdom

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## Impact of Digital Engagement on Weight Loss Outcomes in Obesity Management: A Retrospective Service Evaluation of GLP-1 and Dual GLP-1/GIP Receptor Agonist Therapy in the United Kingdom

Hans Johnson<sup>1, 2\*</sup> MBChB, MRes, MSc; David Huang<sup>1\*</sup> MBBS, BSc; Vivian Liu<sup>1\*</sup> MAS, BSc; Mahmoud Al Ammouri<sup>3\*</sup> MD, MPH; Christopher Jacobs<sup>4\*</sup> MB BChir, MRes, MD (res); Austen El-Osta<sup>3\*</sup> MSc, MPA, PhD

#### **Corresponding Author:**

Hans Johnson MBChB, MRes, MSc Menwell Ltd (t/a Voy) Unit L, Taper Building, 175 Long Lane London GB

#### Abstract

**Background:** Obesity is a global public health challenge. Pharmacological interventions, such as GLP-1 receptor agonists (e.g., semaglutide) and dual GLP-1/GIP receptor agonists (e.g., tirzepatide), have demonstrated significant weight loss. Digital health platforms offering behavioral support may enhance the effectiveness of these medications.

**Objective:** This retrospective service evaluation investigated the impact of engagement with an app-based digital weight loss program on weight loss outcomes among individuals using GLP-1 receptor agonists (semaglutide) and dual GLP-1/GIP receptor agonists (tirzepatide) in the United Kingdom over 5 months.

**Methods:** Data was collected from the Voy weight loss digital health platform between February 2023 and August 2024. Participants were adults aged 18–75 years with a body mass index (BMI) ?30 or ?27.5 kg/m² with presence of obesity-related comorbidities who initiated a weight management program involving semaglutide or tirzepatide. Engagement was defined based on attendance at coaching sessions, frequency of app usage, and regular weight tracking. Participants were categorized as "engaged" or "non-engaged" accordingly. Weight loss outcomes were assessed over a period of up to 5 months. Statistical analyses included chi-square tests, independent t-tests, Kaplan-Meier survival analysis and calculations of Cohen's d for effect sizes.

**Results:** A total of 57,975 participants were included in the analysis, with 31,407 (54.2%) classified as engaged and 26,568 (45.8%) as non-engaged. Engaged participants achieved significantly greater weight loss at each time point. At month 3, engaged participants had a mean weight loss of 9.0% (95% CI: 9.0% to 9.1%) compared to 5.9% (95% CI: 5.9% to 6.0%) in non-engaged participants (P?<?.001), representing a mean difference of 3.1 percentage points (95% CI: 3.1% to 3.1%). Cohen's d effect size of 0.89 indicated a large effect. At month 5, engaged participants had a mean weight loss of 11.53% (95% CI: 11.5% to 11.6%) compared to 8.0% (95% CI: 7.9% to 8.0%) in the non-engaged (P?<?.001). A Cohen's d effect size of 0.56 indicated a moderate effect. Participants using tirzepatide achieved more significant weight loss than those using semaglutide at month 5 (13.9% [95% CI: 13.5% to 14.3%] vs. 9.5% [95% CI: 9.% to 9.71%]; P?<?.001). The proportion of engaged participants achieving ?5%, ?10%, and ?15% weight loss was significantly higher than the non-engaged group at corresponding time points from months 3 to 5 respectively (P?<?.001).

Conclusions: Engagement with a digital weight management platform significantly enhances weight loss outcomes among individuals using GLP-1 receptor agonists. The combination of pharmacotherapy and digital behavioral support offers a promising strategy to promote the supported self-care journey of individuals seeking clinically effective obesity management interventions.

<sup>&</sup>lt;sup>1</sup>Menwell Ltd (t/a Voy) London GB

<sup>&</sup>lt;sup>2</sup>Department of Digital Health and Care, School of Engineering Mathematics and Technology University of Bristol Bristol GB

<sup>&</sup>lt;sup>3</sup>Self-Care Academic Research Unit (SCARU), Department of Primary Care & Public Health, School of Public Health Imperial College London London GB

<sup>&</sup>lt;sup>4</sup>Department of Psychology University of Bath Bath GB

<sup>\*</sup>these authors contributed equally

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## **Original Manuscript**

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#### **Conclusions:**

Engagement with a digital weight management platform significantly enhances weight loss outcomes among individuals using GLP-1 receptor agonists. The combination of pharmacotherapy and digital behavioral support offers a promising strategy to promote the supported self-care journey of individuals seeking clinically effective obesity management interventions.

**Keywords:** obesity; weight loss; semaglutide; tirzepatide; digital health; engagement; behavior; coaching; retrospective study; service evaluation

#### Introduction

#### **Background**

The global obesity epidemic continues to pose a significant challenge to public health systems worldwide [1]. ). Obesity is characterized by excessive fat accumulation that impairs health and is associated with an increased risk of multiple noncommunicable diseases (NCDs), including type 2 diabetes (T2D), cardiovascular diseases and certain cancers [2]. Beyond individual health consequences, this so called "disease of the lifestyle" [3] imposes substantial societal and economic burdens due to elevated healthcare costs and reduced productivity, with a global economic impact estimated at £1.3 trillion [4].

Traditional management strategies for obesity predominantly focus on lifestyle modifications such as dietary changes, increased physical activity and behavioral interventions [1, 5]. While these approaches are fundamental to initiating weight loss and improving health, they often fail to produce long-term results for many individuals [6]. This limitation is partly due to complex physiological adaptations that occur in response to weight loss, including metabolic slowdown and increased appetite, which traditional interventions may not adequately address [7].

Recent advancements include the introduction of pharmacological interventions to promote obesity management. Glucagon-like peptide-1 (GLP-1) receptor agonists (e.g., semaglutide, marketed as Wegovy and Ozempic) [8], and dual GLP-1/gastric inhibitory polypeptide (GIP) receptor agonists like tirzepatide (Mounjaro) [9], represent breakthroughs by targeting physiological pathways that regulate appetite and energy balance. These pharmacotherapies demonstrated the ability to achieve and maintain significant weight loss, ranging from 10% to 20% over two years, serving as promising adjuncts to lifestyle interventions [10, 11].

Notably, the National Institute for Health and Care Excellence (NICE) recognized the significant role of digital tools in supporting weight management and provided guidance on the integration of digital health interventions that offer evidence-based behavioral support to individuals aiming to manage their weight effectively [12]. These tools should be designed to promote sustained weight loss through ongoing engagement, personalized feedback and tailored advice, complementing pharmacological treatments like GLP-1 receptor agonists. The incorporation of such digital platforms aligns with NICE's broader strategy to enhance the accessibility and effectiveness of obesity management programs across diverse populations whilst supporting the weight loss journey of individual self-carers who are overweight but otherwise healthy, as well as patients who need to lose weight to tackle multiformity.

However, significant weight loss is not solely a product of medication use but also depends on sustained behavior change and self-care approaches to promote adherence to treatment protocols. Behavioral factors critically influence the success of weight loss interventions [13]. Supported self-management weight loss programs involving pharmacotherapy, such as the digitally enabled Voy, incorporate behavioral change components and utilize technology to promote end-user engagement. By integrating behavior change theories like Social Cognitive Theory [14] and facilitating self-monitoring, goal setting and feedback mechanisms, these platforms encourage patients to actively manage their health and wellbeing journey in the community and other settings [15] [16]. Behavioral activation complements weight loss interventions by helping individuals identify and engage in positive, goal-oriented activities that align with their health objectives. It emphasizes breaking the cycle of avoidance or inactivity often associated with obesity, replacing these behaviors with structured, rewarding actions like regular physical activity, healthy eating, and consistent self-monitoring [17]. By focusing on small, achievable steps, behavioral activation builds motivation and self-efficacy, which are crucial for sustained adherence to treatment protocols. When combined with digital tools, such as goal-setting features and real-time feedback, this approach empowers individuals to take ownership of their weight management journey, enhancing both engagement and long-term outcomes [18]. Multicomponent interventions are attractive because they address both the physiological and behavioral aspects of obesity. This approach potentially enhances the effectiveness of pharmacological treatments through ongoing support and education using real-time microlearning approaches to promote individual self-care capability [19], as well as through supporting the adoption of health-seeking self-care behaviors using nudges and personalized insights delivered using an accessible digital platform [20, 21].

Clinically significant weight loss is typically defined as a reduction of at least 5% of initial body weight [22], which is associated with improvements in obesity-related comorbidities such as hypertension, dyslipidemia, and insulin resistance [23]. Achieving weight loss of 10% or more is linked to additional health benefits, including enhanced glycemic control, decreased need for diabetic medications, and reduced risk of cardiovascular events [24]. Weight loss of 15% or greater can lead to substantial clinical improvements, such as remission of type 2 diabetes and significant reductions in cardiovascular risk factors [25]; ultimately interventions in obesity management that aim to reduce weight should ideally also reach thresholds that confer meaningful health benefits.

Digitally supported self-care approaches [26] and supported self-management are integral to lifestyle medicine [27]. Despite the rapid adoption of digital health technologies, there remains a substantial gap in understanding how these digital interventions can complement pharmacological treatments in obesity for improved clinical outcomes compared to pharmacological treatment alone. Randomized controlled trials have examined eHealth platforms with behavioral change coaching for weight loss [28]. However, existing literature is limited, as the integration of digital platforms with pharmacological therapy for weight loss has yet to be extensively researched due to its novelty [29-31].

#### **Objectives**

The aim of this study was to address this gap by exploring the impact of engagement with appbased digital weight loss programs that combine pharmacotherapy using GLP-1 receptor agonists semaglutide and the dual GLP-1/GIP receptor agonist tirzepatide. Considering that the ideal focus is on achieving and sustaining clinically significant weight loss through digital engagement support over specified durations, this study sought to provide valuable insights

into the potential and limitations of digital health strategies in enhancing the efficacy of obesity management.

#### **Methods**

#### **Study Design and Setting**

This retrospective 2-arm study was conducted to assess the effectiveness of GLP-1 receptor agonists. Specifically semaglutide (marketed as Wegovy and Ozempic) and the dual GLP-1/GIP receptor agonist tirzepatide (marketed as Mounjaro) within a digitally delivered weight management program. The evaluation utilized data extracted from the Voy digital health platforms, which provided remote support through digital tools, including direct delivery of pharmacotherapy.

The study period spanned February 2023 to August 2024, including up to 5 months of data for participants using both semaglutide and tirzepatide. The selection of 5-month follow-up duration was based on data availability, representing a cross-sectional cohort of individuals who were recently onboarded to the platform for the weight loss service, allowing for a consistent assessment of early treatment outcomes across all participants in the initial 5 months where significant weight loss is typically observed [9].

#### **Procedure**

Participants enrolled in the Voy digital health platform's weight management program, which integrates GLP-1 receptor and GLP-1/GIP receptor agonist pharmacotherapies with digital behavioral support to enhance weight loss outcomes (see Figure 1. flow chart). Upon enrollment, participants underwent an initial assessment to confirm eligibility, including verification of age, BMI, and absence of exclusion criteria as outlined in *eligibility criteria* below. Participants also completed online asynchronous consultations, which included medical suitability checks, identity verification through photo ID, and submission of any required documentation. Baseline demographic information, medical history and lifestyle factors were requested at the initiation of the program.

Eligible participants received prescriptions for either semaglutide or tirzepatide based on clinical considerations. Medications were delivered directly to participants, with comprehensive medication management provided through the platform. Instructional materials ensured correct medication administration, and participants had unlimited consultations with guidance and support from a team of clinicians and coaches.

Participants attended group onboarding sessions and were offered fortnightly coaching to enhance engagement and adherence. Coaches were trained based on principles from Social Cognitive Theory, Self-Determination Theory, the Transtheoretical Model, and the Theory of Planned Behavior [14, 32-34]. These techniques focused on fostering intrinsic motivation, goal setting, and problem-solving to promote sustainable lifestyle changes tailored to participants' individual progress and challenges [21]. Participants were encouraged to actively engage with the App's dynamic educational content, which adapted based on their engagement with specific topics, including nutrition, physical activity, and lifestyle factors impacting weight management. The educational content aimed to enhance self-efficacy and equip participants with practical skills to sustain weight loss over time [35].

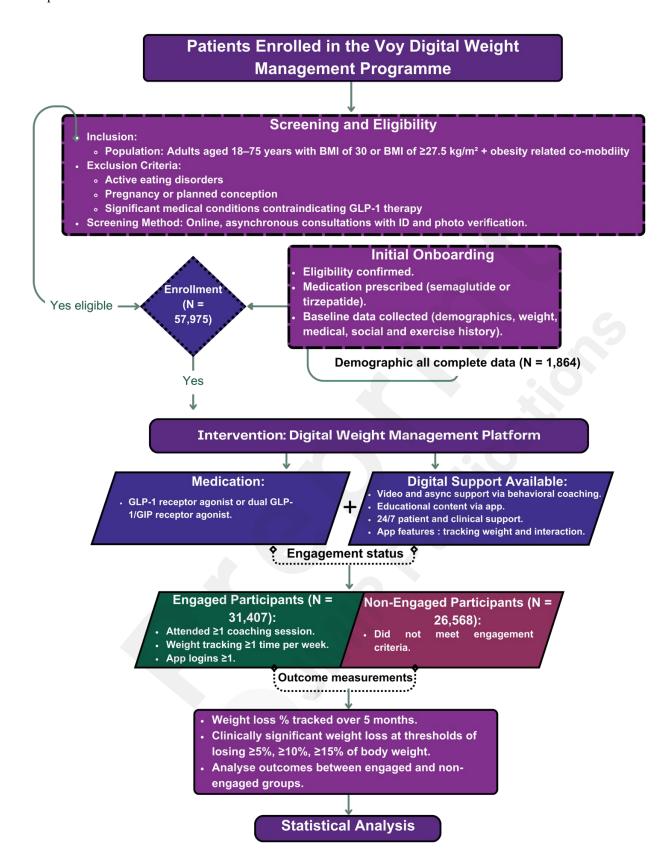


Figure 1. Flowchart illustrating the critical steps of the retrospective analysis conducted for the Voy digital weight management programme.

#### **Participants**

Participants were adult residents of the United Kingdom, aged between 18 and 75 years, with a body mass index (BMI) of 30 kg/m $^2$  or higher, or >27.5 kg/m $^2$ , with presence of obesity related comorbidities. All participants initiated a weight management program involving either semaglutide or tirzepatide.

#### **Eligibility Criteria**

Eligibility required access to a smartphone or tablet to engage with the digital health platform. Exclusion criteria included history of self-reported eating disorders (e.g., anorexia nervosa, bulimia nervosa), pregnancy or active attempts to conceive, known allergies or hypersensitivity to any components of the prescribed medications, and severe medical conditions such as a personal or family history of medullary thyroid carcinoma, multiple endocrine neoplasia syndrome type 2, significant hepatic impairment, renal impairment requiring dialysis, uncontrolled cardiovascular diseases, or severe gastrointestinal disorders (e.g., gastroparesis, pancreatitis). Participants who met the eligibility criteria were identified through the digital health platforms' databases. All eligible individuals who initiated the program within the study period were included, resulting in a sample reflective of real-world clinical practice.

#### **Data Availability**

Engagement metrics and weight loss outcomes were available for the entire sample of 57,975 participants who used the digital platform during the study period. Complete demographic information (e.g., age, sex, ethnicity) was available for a subset of 1,864 participants who enrolled during the initial phase of the service. This subset represents the early adopters for whom detailed baseline data were systematically collected to inform program development and initial evaluations.

#### **Defining Engagement and Outcome**

The primary outcome measured in this study was the degree of weight loss measured in percentage weight change. Engagement with the digital platform's program was evaluated as an exposure variable, with levels of engagement assessed for their impact on the outcome of weight loss.

Participants were categorized into "engaged" or "non-engaged" groups based on their interaction with the following components of the program: (i) attendance at weight coaching sessions, (ii) regularity of weight tracking, and (iii) frequency of app usage. The categorization of engagement was determined with input from the clinical team delivering the intervention.

Participants were classed as engaged if they met any number of the below criteria during the study: (i) attended at least one weight coaching session, (ii) tracked weight at least once per week, and/or (iii) logged into the App at least once. Otherwise, participants were classified as non-engaged.

#### **Variables**

The primary outcome of the study was the percentage weight change from baseline, calculated as the aggregated average of each individual's weekly weight change converted to a monthly average and then pooled across all participants. This outcome included the effect of weight loss medication use, reaching clinically significant weight loss thresholds defined as loss of  $\geq 5\%$ ,  $\geq 10\%$ , and  $\geq 15\%$  of baseline weight. Engagement with the digital platform was assessed as an

exposure variable, whereas engagement level was measured by quantifying components such as attendance at coaching sessions, tracking weight within the app and logging into the app. Other exposure variables included the type of medication used, categorized as semaglutide or tirzepatide.

#### **Statistical Analysis**

Descriptive statistics were used to summarize baseline characteristics, with means and standard deviations (SD) for continuous variables and frequencies and percentages for categorical variables. Chi-square tests were used to compare the proportion of participants achieving clinically significant weight loss ( $\geq 5\%$ ,  $\geq 10\%$ , and  $\geq 15\%$  of baseline weight) between the engaged and non-engaged groups. In cases where small sample sizes or low expected frequencies were present (i.e. n <5), Fisher's exact test was used. Odds ratios (ORs) and 95% confidence intervals (CIs) were calculated to quantify the strength of the association between engagement and weight loss success through 2x2 contingency tables.

To assess the time-to-event data for achieving clinically significant weight loss, Kaplan-Meier survival analysis was performed, comparing the time to reach  $\geq 5\%$ ,  $\geq 10\%$ , and  $\geq 15\%$  weight loss between the engaged and non-engaged groups. The log-rank test was used to evaluate the statistical significance of differences between the survival curves of the two groups. This approach allowed us to visualize the cumulative incidence of weight loss over time and assess the role of engagement in accelerating weight loss success.

An independent sample t-test was used to compare the mean weight change between the engaged and non-engaged groups. Cohen's d was calculated to determine the effect size, providing insight into the magnitude of differences between groups. Prior to conducting the t test, assumptions of normality and homogeneity of variances were assessed using the Shapiro–Wilk test. When assumptions were violated, a nonparametric alternative, the Mann–Whitney U test, was considered.

All statistical analyses were performed using R (version 4.3.1) and its statistical packages. A significance level of P < .05 was set for all statistical tests.

#### **Sample Size Calculation**

A sample size calculation was performed to estimate the number of participants required to detect a significant difference in weight loss between engaged and non-engaged groups. Using an estimated effect size of a 15% difference in the proportion of participants achieving  $\geq 10\%$  weight loss between groups, with an expected 30% of engaged participants and 15% of non-engaged participants achieving this outcome, a minimum of 118 participants per group was calculated to provide 80% power at a 5% significance level. To account for attrition, a target sample size of 250 participants per group was set.

Baseline demographic information, medical history and lifestyle factors were requested at the initiation of the program. Data concerning medication adherence were monitored through self-reports and prompted reminders. All data were anonymized upon extraction to ensure confidentiality in compliance with the General Data Protection Regulation (GDPR).

#### **Bias and Missing Data**

Self-reported weight measurements could introduce reporting bias. To mitigate this, participants were encouraged to provide accurate measurements through regular reminders

and had the option to upload progress photographs, enhancing data validity. Additionally, data validation checks were performed to identify and address implausible values. Selection bias was minimized by including all eligible participants who initiated the program within the study period, ensuring the sample was representative of the population utilizing these services.

#### **Ethical Considerations**

Imperial College Research Ethics Committee (REC) provided a favorable opinion for this study (ICREC#7363051). As this evaluation utilized anonymized data collected during routine care, formal ethical approval from the National Health Service (NHS) REC was not required under NHS standards for service evaluations. The study adhered to the principles outlined in the Declaration of Helsinki. Participants provided informed consent for their anonymized data to be used for research and service improvement purposes upon enrollment in the program. Data protection and confidentiality were strictly maintained in compliance with GDPR regulations.

#### **Adherence to STROBE Guidelines**

To improve the quality of the reporting, this study adhered to the STROBE (Strengthening the Reporting of Observational Studies in Epidemiology) guidelines [36] (See Supplementary. File 1).

#### Results

#### **Participant Characteristics**

A total of 57,975 participants were included in the engagement and outcome analyses. Among these, 31,407 (54.2%) were classified as engaged and 26,568 (45.8%) as non-engaged at baseline (month 0). Due to changes in data collection practices over time and staggered onboarding during the study period, the number of participants reporting weight data decreased at subsequent time points, comprehensive demographic and clinical characteristics were available for a subset of 1,864 participants who enrolled at the inception of the service. The characteristics of this subset are presented in **Table 1**.

Table 1. Baseline characteristics of participants with available data (N = 1,864).

Characteristic	Value
Age, mean (SD), years	45.2 (± 10.5)
Sex, n (%)	<b>○</b>
	1,592
Female	(85.4%)
Male	272 (14.6%)
BMI, mean (SD), kg/m <sup>2</sup>	34.9 (±4.9)
Weight, mean (SD), kg	96.2 (±20.9)
Ethnicity, n (%)	
	1,439
White British/Irish	(77.2%)
White Other	199 (10.7%)
Asian Indian	39 (2.1%)

Asian Dalristani	01 (1 10/)
Asian Pakistani	21 (1.1%)
Black African	32 (1.7%)
Other Ethnic Groups	134 (7.2%)
Comorbidities, n (%)	20 (4 204)
Type 2 Diabetes	80 (4.3%)
High Blood Pressure	289 (15.5%)
High Cholesterol	193 (10.4%)
Osteoarthritis	123 (6.6%)
Chronic Back Pain	231 (12.4%)
Depression	512 (27.5%)
Lifestyle Factors, n (%)	
Smoking Status	
Never Smoked	945 (50.7%)
Ex-smoker	433 (23.2%)
Current Smoker	108 (5.8%)
Not Answered	378 (20.3%)
Alcohol	
Consumption/weekly	
None (0 units)	765 (41.1%)
1–6 units	235 (12.6%)
7+ units	864 (46.3%)
Dietary Habits	
	1,238
Balanced Diet	(67.5%)
High in Animal Products	293 (16.0%)
Vegetarian/Vegan	132 (7.1%)
Not Answered	175 (9.4%)
Physical Activity	
Daily	180 (9.7%)
Regularly (3–4/week)	626 (33.6%)
Occasionally (1–2/week)	702 (37.7%)
Rarely/Never	356 (19.1%)
Family History of Obesity,	
n (%)	988 (53.1%)
Medications, n (%)	
Blood Pressure Meds	211 (11.3%)
Cholesterol Meds	182 (9.8%)
Diabetes Meds	136 (7.3%)
Thyroid Meds	97 (5.2%)
Antidepressants	247 (13.2%)

#### **Engagement Levels**

At baseline (month 0), among the 57,975 participants who tracked their weight, 31,407

(54.2%) were engaged, and 26,568 (45.8%) were non-engaged. Engagement levels decreased over time, with 3,622 (6.3%) engaged participants remaining at month 5 (Table 2.).

Table 2. Mean percentage of weight loss by engagement in participants taking medication and using the digital platform. All weight loss values are mean aggregates and are in a negative

value. \*P-values derived from independent t-test.

Month from Medica tion Start	Engag ement	Mean % Weight Lost (95% CI)	Absolute % Point Difference	<i>P</i> - Value*	Effect size (d)	Relative % Differe nce	Numbe r of Patient s	Total Patient s
0	No	N/A					26568	
0	Yes	N/A	0.00	N/A	0.00	N/A	31407	57975
1	No	2.87 (2.86 - 2.88)					1429	
1	Yes	3.84 (3.83 - 3.85)	0.97	<.001	0.179	33.8	21877	23306
2	No	5.03 (5.01 - 5.05)					1848	
2	Yes	6.90 (6.89 - 6.91)	1.88	<.001	0.291	37.4	12978	14826
3	No	5.93 (5.90 - 5.96)				20	1691	
3	Yes	9.04 (9.02 - 9.06)	3.11	<.001	0.891	52.4	8429	10120
4	No	6.99 (6.95 - 7.03)					1331	
4	Yes	10.67 (10.64 - 10.70)	3.68	<.001	0.616	52.7	5495	6826
5	No	7.97 (7.93 - 8.01)	3.53	<.001	0.560	44.3	1377	4999

#### **Weight Loss Outcomes**

#### Weight Loss by Engagement Status

Engaged participants consistently achieved greater weight loss compared to non-engaged participants at each time point (Figure 2). At month 1, engaged participants experienced a mean weight loss of 3.8% (95% CI: 3.9% to 3.8%), whereas non-engaged participants had a mean weight loss of 2.9% (95% CI: 2.9% to -2.9%), a significant difference (P < .001). This trend persisted over subsequent months. By month 5, engaged participants had a mean weight loss of 11.5% (95% CI: 11.6% to 11.5%), while non-engaged participants had a mean weight loss of 8.0% (95% CI: 8.0% to 7.9%), a significant difference of 3.56 percentage points (P < .001).

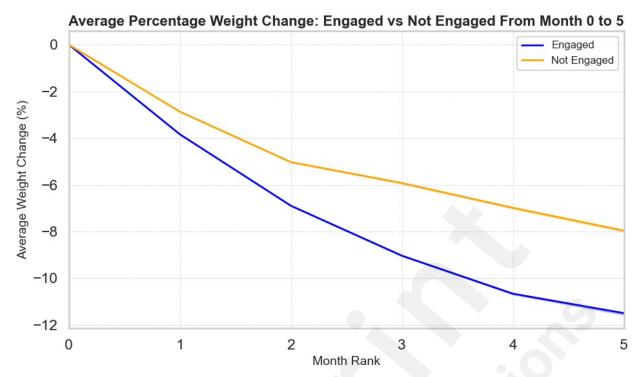


Figure 2. Average weight loss (Kg) trajectory over time by engagement status.

#### Weight Loss by Medication Type

Participants were prescribed either semaglutide or tirzepatide (Mounjaro). Weight loss trajectories differed significantly between the two medication groups over the study period (Figure 3.; Appendix 1. Table 6.). At month 5, participants using tirzepatide achieved a mean weight loss of 13.9% (95% CI: 14.3% to 13.5%), significantly greater than the 9.5% (95% CI: 9.7% to 9.2%) observed in semaglutide users (P < .001).

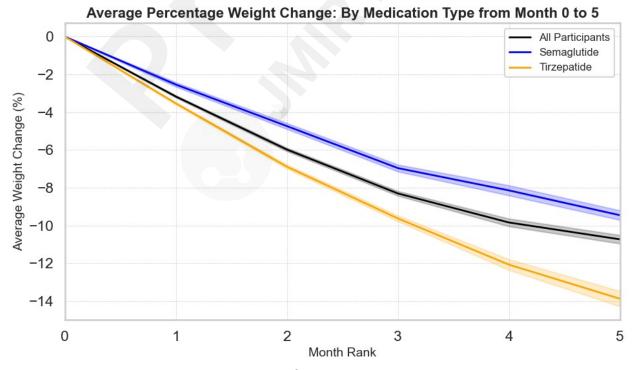


Figure 3. Depicts the mean percentage of weight loss over time by medication type. 'All

participants' shows the combined weight loss trajectory for both tirzepatide and semaglutide users.

#### **Proportion Achieving Clinically Significant Weight Loss**

The proportion of participants achieving clinically significant weight loss thresholds ( $\geq$ 5%,  $\geq$ 10%, and  $\geq$ 15% of baseline weight) was higher among engaged participants compared to non-engaged participants. At the  $\geq$ 5% weight loss threshold, engaged participants had significantly higher odds of achieving this goal compared to non-engaged participants at all assessed time points. **Table 3** Presents the detailed odds ratios for achieving  $\geq$ 5% weight loss over time.

Table 3. Odds Ratios for Achieving  $\geq 5\%$  Weight Loss by Engagement Status. \*P-value derived

from Chi-squared test.

Mont	Engaged Achieved	Engaged Did Not Achieve	Non- Engaged Achieved	Non- Engaged Did Not Achieve	Odds	OFW OF	D.V. L *
h	Goal (n)	Goal (n)	Goal (n)	Goal (n)	Ratio	95% CI	<i>P</i> -Value*
1	3,083	17,332	104	1,066	1.82	1.49 - 2.24	<.001
2	6,386	6,678	140	612	4.18	3.47-5.04	<.001
3	5,454	3,522	136	352	4.01	3.27-4.91	<.001
4	3,850	2,198	99	214	3.79	2.97-4.83	<.001
5	2,664	1,705	69	117	2.65	1.96-3.59	<.001

For the  $\geq 10\%$  weight loss threshold, significant differences emerged from month 2 onward. Table 4 provides the odds ratios for achieving  $\geq 10\%$  weight loss over time.

At the  $\geq 15\%$  weight loss threshold, significant differences were observed at months 3 and 4. Table 5 details the odds ratios for achieving  $\geq 15\%$  weight loss over time.

Table 4. Odds Ratios for Achieving ≥10% Weight Loss by Engagement Status. \**P*-value derived

from Chi-squared test.

Month	Engaged Achieved Goal (n)	Engaged Did Not Achieve Goal (n)	Non- Engaged Achieved Goal (n)	Non- Engaged Did Not Achieve Goal (n)	Odds Ratio	95% CI	<i>P-</i> Value*
1	331	20,084	25	1,145	0.75	0.50-1.14	0.22
2	1,246	11,818	26	726	2.94	1.98-4.37	<.001
3	2,378	6,598	51	437	3.09	2.30-4.14	<.001
4	2,217	3,831	39	274	4.07	2.90-5.71	<.001
5	1,722	2,647	34	152	2.91	2.00-4.24	<.001

Table 5. Odds Ratios for Achieving  $\geq 15\%$  Weight Loss by Engagement Status. \**P*-value derived from Chi-squared test.

Mont	Engaged	Engaged Did	Non-	Non-	Odds	95% CI	<i>P-</i> Value*

h	Achieved Goal (n)	Not Achieve Goal (n)	Engaged Achieved Goal (n)	Engaged Did Not Achieve Goal (n)	Ratio		
1	93	20,322	12	1,158	0.44	0.24-0.81	.01
2	176	12,888	11	741	0.92	0.50-1.70	0.92
3	509	8,467	9	479	3.20	1.64-6.22	<.001
4	793	5,255	14	299	3.22	1.88-5.54	<.001
5	752	3,617	20	166	1.73	1.08-2.76	.03

#### Time to Achieve Weight Loss Thresholds

Kaplan-Meier analyses (Figures 4–6) revealed significant differences in the proportions of individuals achieving 5%, 10%, and 15% weight loss over 5 months between engaged and nonengaged participants. For example, at month 4, the proportion not achieving the 5% threshold was 49.3% for engaged participants compared to 69.8% for non-engaged participants (P < .001). Similarly, by month 5, 59.9% of engaged participants had yet to achieve the 10% threshold compared to 81.5% of non-engaged participants (P < .001). For the 15% threshold, while differences were less pronounced, 86.5% of engaged participants had not reached this target by month 5, compared to 93.5% of non-engaged participants (P = .017). These findings underscore the role of engagement in facilitating weight loss, with the effect diminishing at higher thresholds.

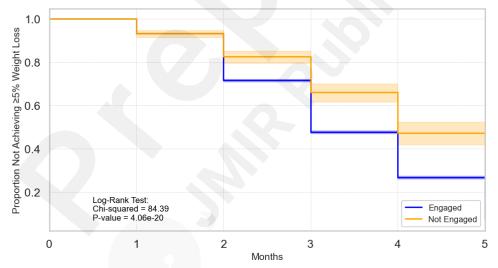


Figure 4. Kaplan–Meier survival curves for time to achieve ≥5% weight loss by engagement status.



Figure 5. Kaplan–Meier survival curves for time to achieve ≥10% weight loss by engagement status.

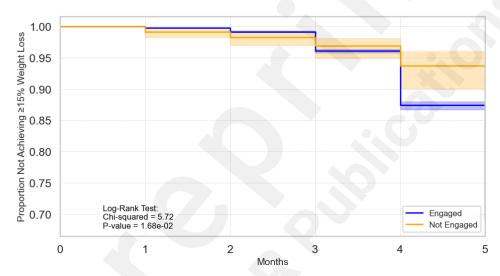


Figure 6. Illustrates the Kaplan–Meier survival curves for time to achieve ≥15% weight loss by engagement status.

#### **Discussion**

#### **Principal Results**

This study provided compelling evidence that engagement with digital health platforms significantly enhances weight loss outcomes among individuals undergoing pharmacotherapy with GLP-1 receptor agonists for obesity management. Engaged participants achieved greater mean weight loss over time and reached clinically significant weight loss thresholds more rapidly than their non-engaged counterparts. The superior efficacy of tirzepatide over semaglutide observed in this real-world study aligns with findings from recent clinical trials reinforcing the potential of dual agonist therapies in obesity management [37].

#### Limitations

The principal limitation of this study was the potential selection bias, as participants who chose

to engage with the digital platform may inherently possess higher motivation levels or greater health literacy. This could confound the relationship between engagement and weight loss outcomes. Although we sought to minimize this selection bias by including all eligible participants, unmeasured confounding must be considered. The discrepancy in available demographic data and the reduction in participant numbers over time may also introduce attrition bias. We also acknowledge that missing data on key variables limits the ability to adjust for confounders and affects the representativeness of the sample.

The discrepancy in demographic data collection reflects adaptations made during program scaling to improve accessibility. Initially, comprehensive data were collected; however, this was streamlined to simplify onboarding and enhance user experience, prioritizing essential data for engagement analysis. While this approach limits detailed subgroup analysis, the broader dataset ensures robust insights into program effectiveness. The collection of data and availability of participant information may account for reducing participant numbers at time intervals. Also, participants may have chosen to no longer engage with the provider. This was retrospective data, and missing observations are more common [38].

Reliance on self-reported weight data also introduces the potential for reporting bias, as participants may underreport or overreport their weight due to social desirability or recall bias. The provision of options to upload progress photographs may mitigate this but does not eliminate the issue. Crucially, the absence of a randomized control group limits the ability to establish causality definitively. Observed associations may also be influenced by unmeasured confounding variables or external factors. Pertinently, the maximum follow-up period of 5 months may not capture long-term weight maintenance or the sustainability of engagement with the digital platform. Given that weight regain is a common challenge in obesity management, longer-term studies are needed to assess enduring effects.

The study population was predominantly female and of White British/Irish ethnicity, which also may limit the generalizability of the findings to more diverse populations as cultural, socioeconomic and gender differences may influence engagement and weight loss outcomes. Additionally, whereas only a subset of 1,864 participants was available for analysis, we acknowledge that this subset may not likely be representative of the total population.

We also acknowledge that the requirement for smartphone or tablet access may exclude individuals from lower socioeconomic backgrounds or older adults less comfortable with technology, potentially exacerbating health disparities.

Building on the insights from this study, future research should conduct RCTs comparing pharmacotherapy with and without digital behavioral support to establish causality and quantify the added benefit of digital engagement. Qualitative studies investigating user experiences, barriers to engagement and facilitators of sustained use can inform the design of more effective digital interventions. Longitudinal studies with extended follow-up periods are needed to evaluate the sustainability of weight loss and the impact of ongoing engagement on weight maintenance and metabolic health. Efforts to collect comprehensive demographic and clinical data for all participants will improve the ability to adjust for confounding factors and understand the differential effects across subgroups.

Research should focus on strategies to streamline timely access to supported self-management technologies, including pharmacotherapy, cognitive behavior therapy approaches and digital

self-care interventions among diverse populations, including those with limited digital literacy, to ensure equitable benefits and reduce health disparities. Incorporating microdata from wearables, biomarker assessments, and validated scales for psychological measures, quality of life and individual self-care capability can enhance the accuracy of outcomes and provide a more holistic evaluation of health improvements.

#### **Comparison with Prior Work**

The integration of supported self-care interventions involving pharmacotherapy into obesity management represents a major shift that can help address the various challenges associated with weight loss and maintenance. Digital platforms offer scalable, personalized and accessible supported self-management solutions that transcend traditional barriers to healthcare delivery [39]. In this example, the digital platforms facilitated self-monitoring, behavioral coaching and personalized feedback components that are critical to promoting sustained behavior change. The theoretical foundations of digital interventions are grounded in established behavior change models such as the Transtheoretical Model and Self-Determination Theory [40], emphasizing self-efficacy, intrinsic motivation and readiness to change [41]. By providing real-time feedback and reinforcing positive behaviors, digital platforms can strengthen self-regulatory processes essential for weight management [42].

The increasing pervasiveness of mobile health technologies facilitates continuous engagement and data collection [43] and presents an unprecedented opportunity for self-driven healthcare approaches to deliver interventions at scale and reach populations traditionally underserved or facing barriers to in-person care [26]. The enhanced weight loss observed among engaged participants highlights the potential of digital platforms to augment traditional treatment modalities.

Several synergistic mechanisms likely contributed to the enhanced weight loss observed among engaged participants. Enhanced self-monitoring and accountability are fundamental, as frequent tracking of weight and behaviors is a consistent predictor of weight loss success [44]. Digital self-care platforms simplify this process through user-friendly interfaces and automated reminders, reducing the burden of manual tracking and increasing adherence (37). Tailored interventions have been shown to be more effective than generic advice in promoting behavior change for weight loss [45]. This is likely because personalized behavioral support provided through coaching and tailored feedback also addresses individual barriers and facilitators to weight loss, further enhancing motivation and self-efficacy [46]. The principles of behavioral activation with a focus on action following analysis of behaviors enables a structured problemsolving approach that provides a treatment foci related to change [47]. This incorporation of evidence-based behavior change techniques, such as goal setting, action planning and problemsolving, is associated with improved weight loss outcomes [48]. Features enabling interaction with peers or health professionals provide social support, which is a critical determinant of weight management success [49], whereas social comparison and normative influence may further motivate adherence to weight loss plans [50]. Regular engagement with the app may also facilitate the formation of automatic healthy behaviors through repetition and reinforcement, as suggested by Habit Formation Theory [51]. Combining behavioral interventions with pharmacotherapy may produce a synergistic effect, enhancing the biological mechanisms of weight loss medications through improved adherence and lifestyle modifications [52].

Our findings align with and extend existing literature on integrating digital interventions with pharmacological treatments for obesity. Previous studies have demonstrated that digital behavioral interventions can enhance the effectiveness of weight loss medications [53]. For example, a meta-analysis by Beleigoli et al. [54] found that web-based interventions resulted in modest but significant additional weight loss compared to standard care. The superior weight loss outcomes associated with tirzepatide in our study are also consistent with clinical trials demonstrating its efficacy over other GLP-1 receptor agonists [55, 56]. We acknowledge that the dual action on GIP and GLP-1 receptors may confer additive or synergistic effects on glycemic control and appetite regulation, leading to greater weight reductions [57].

#### **Conclusions**

This study provided compelling evidence that structured engagement with a digital self-management platform significantly enhances weight loss outcomes by as much as 53% at month four in engaged individuals undergoing pharmacotherapy for obesity. The integration of digital behavioral support with pharmacological treatments represents a synergistic approach that addresses both the biological and behavioral dimensions of obesity. By facilitating self-monitoring, providing personalized feedback and using nudge and gamification to promote sustained engagement, digital self-management tools can amplify the effectiveness of pharmacotherapies. As the prevalence of obesity continues to rise globally, innovative self-driven healthcare solutions that leverage technology to support behavior change are needed to scale benefits to individuals with non-communicable diseases, and so called "diseases of the lifestyle".

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#### **Author Contributions**

All authors contributed to the conception and design of the study. HJ performed the data analysis and interpretation of the results, along with MA (interpretation). VL and HJ ran the statistical analysis, with assessment of analysis by MA. HJ, DH,VL, CJ, and AO drafted the manuscript, and all authors critically reviewed and approved the final version for publication

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#### Conflicts of Interest

The authors HJ, DH, and VL are members within the organization Voy, Menwell<sup>IM</sup>; HJ is the clinical researcher, DH is the innovation director, and VL is the research lead. The authors AO, MA and CJ declare that they have no conflicts of interest relevant to the content of this article.

#### **Abbreviations**

BMI: body mass index

CI: confidence interval

GDPR: General Data Protection Regulation

GIP: gastric inhibitory polypeptide GLP-1: glucagon-like peptide-1 NHS: National Health Service

OR: odds ratio

RCT: randomized controlled trial

SD: standard deviation

STROBE: Strengthening the Reporting of Observational Studies in Epidemiology

#### Appendix 1

Table 6. The table provides a detailed summary of weight change data across six time points (Month Rank 0 to 5) for three groups: All participants, those on Semaglutide, and those on Mounjaro. \*P-value derived from independent t-test.

Month Rank	Avg Weight Change (%) - All (95% CI)	Avg Weight Change (%) - Semaglutide (95% CI)	Avg Weight Change (%) - Tirzepatide (95% CI)	P-Value* (Semaglutide vs Tirzepatide)
0	0 (0-0)	0 (0-0)	0 (0-0)	N/A
1	-3.17 (-3.233.11)	-2.54 (-2.662.42)	-3.54 (-3.603.47)	<.001
2	-5.98 (-6.075.89)	-4.74 (-4.894.58)	-6.89 (-6.996.78)	<.001
3	-8.31 (-8.438.18)	-6.96 (-7.146.78)	-9.63 (-9.789.48)	<.001
4	-9.84 (-10.05 9.64)	-8.14 (-8.417.88)	-12.08 (-12.36 11.79)	<.001
5	-10.74 (-10.9610.51)	-9.46 (-9.719.21)	-13.88 (-14.29 13.48)	<.001

#### **Supplementary File 1**

#### S.1 Adherence to STROBE Guidelines

This study adhered to the STROBE (Strengthening the Reporting of Observational Studies in Epidemiology) guidelines. Attached with paper.

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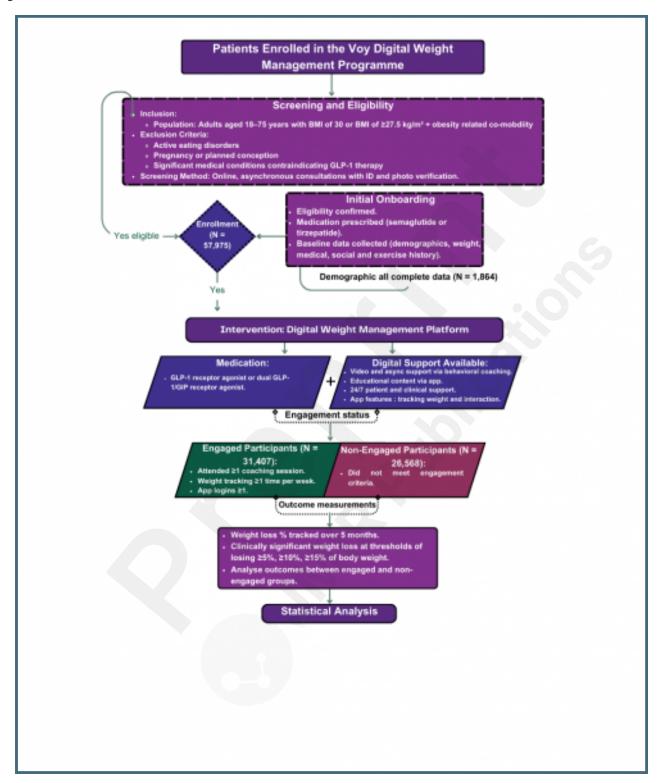
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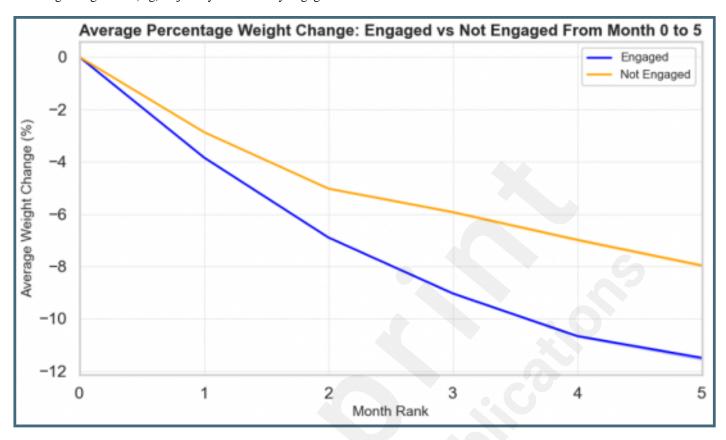
## **Supplementary Files**

## **Figures**

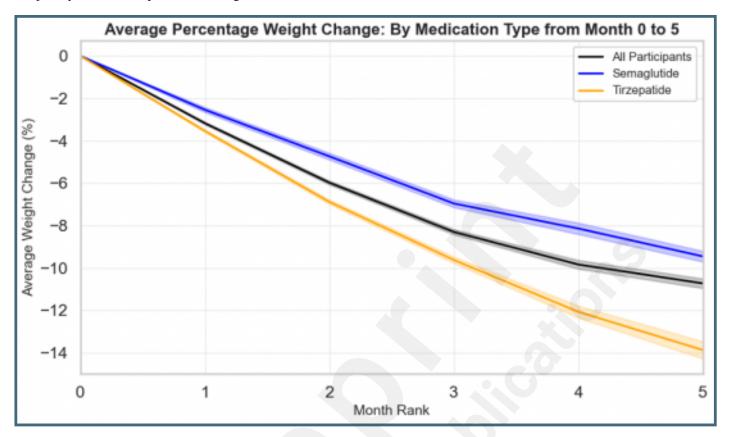
Flowchart illustrating the critical steps of the retrospective analysis conducted for the Voy digital weight management programme.



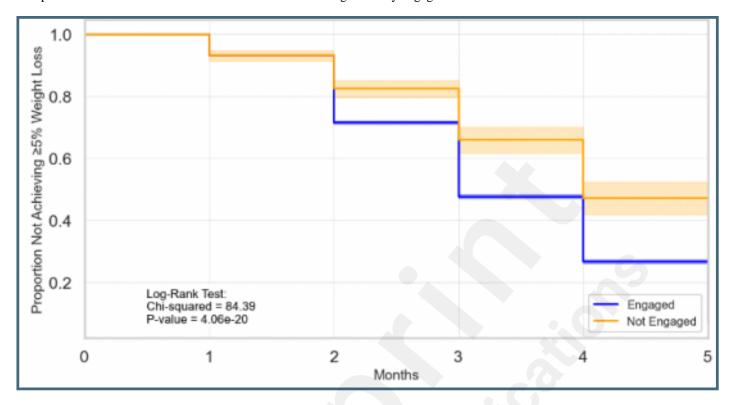
Average weight loss (Kg) trajectory over time by engagement status.



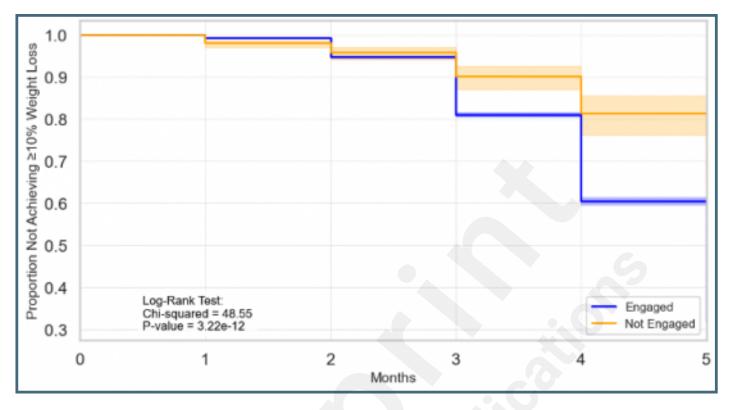
Depicts the mean percentage of weight loss over time by medication type. 'All participants' shows the combined weight loss trajectory for both tirzepatide and semaglutide users.



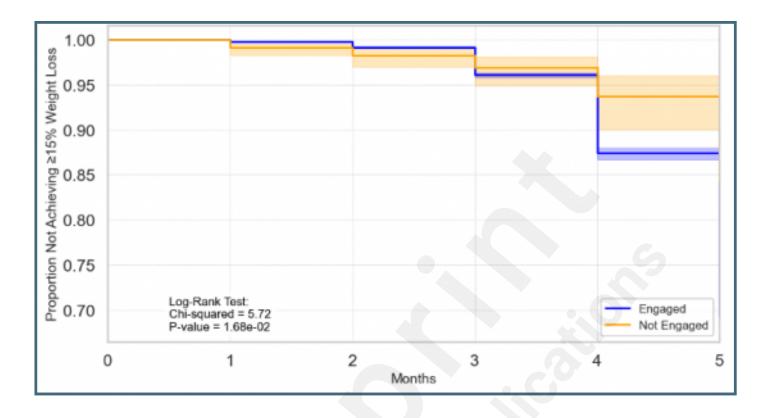
Kaplan-Meier survival curves for time to achieve ?5% weight loss by engagement status.



Kaplan-Meier survival curves for time to achieve ?10% weight loss by engagement status.



Illustrates the Kaplan-Meier survival curves for time to achieve ?15% weight loss by engagement status.



### **CONSORT** (or other) checklists

Adherence to STROBE Guidelines This study adhered to the STROBE (Strengthening the Reporting of Observational Studies in Epidemiology) guidelines. Attached with paper.

URL: http://asset.jmir.pub/assets/17f38765e4ee5ada744de778963280dd.pdf