

# Characteristics of Patients Initiating Glucagon-like Peptide-1 (GLP-1) Receptor Agonists (RAs) for Cardiometabolic Risk Reduction in a Medicare Population

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## Background and Rationale

- Glucagon-like Peptide-1 (GLP-1) Receptor Agonists (RAs) have been approved for Type 2 diabetes (T2D), and obesity.
- More recently, in March 2024, Wegovy® (semaglutide) became the first GLP-1 RA approved for cardiovascular risk reduction, followed by Ozempic® (semaglutide) approval in January 2025 for reduction in the risk of worsening kidney disease and cardiovascular death in adults with T2D and chronic kidney disease.
- However, GLP-1 RA trends across these indications have not been well characterized among Medicare beneficiaries.

**Objective:** to describe clinical characteristics, treatment patterns, and outcomes of GLP-1 RA initiators in the US Medicare 100% Fee-for-Service (FFS) population.

## Methods



### Study Design

- We conducted a retrospective observational cohort study of patients initiating GLP-1 RAs using the 100% Medicare FFS claims database.
- Data Source:** This study uses data from the 100% Medicare FFS database and pharmacy data. The Medicare FFS is a traditional fee-for-service health plan with two parts: Part A [Hospital Insurance] and Part B [Medical Insurance]). Part B insurance contains information related to inpatient, outpatient, and office visits.



### Eligibility Criteria

- All patients initiating GLP-1 RAs between January 01, 2018–June 30, 2024.
  - At least 18 years of age at index date.
  - At least 6 months of continuous health plan and pharmacy enrollment (Part A, Part, B, & Part D) prior to the index date.
- Outcomes**
- Real-world overall survival (rwOS), defined as death in the follow-up period.
  - Myocardial infarction (MI), defined as the presence of ICD-10-CM code of I21.XX for acute myocardial infarction in the follow-up period.
  - Discontinuation was defined as a gap of >90 days between one claim/pharmacy fill date plus days supply and the subsequent fill date.



### Analysis

- Index date:** date of GLP-1 RA initiation within study period.
- Baseline and clinical characteristics within 6 months prior to index date were described.
- Utilization was described by generic drug over years.
- Outcomes (rwOS, MI) were presented overall and by generic drug.
- Time to event analysis**
- Patients indexed between 2018-2023 were included to allow for the opportunity of ≥6 months of follow-up.
- rwOS:** time from index date to earliest of date of death, or end of enrollment, discontinuation of any GLP-1 RA drug, or end of study period (censoring criteria).
- MI:** time from index date to earliest of MI, or date of death, end of enrollment, discontinuation of any GLP-1 RA drug, or end of study period (censoring criteria).
- Additional censoring criteria for by-drug analysis included discontinuation of the specific generic, or date of switch to different generic drug.

## Key Findings

- Median age of patients initiating GLP-1 RA was 69 (IQR: 65-74) and more than half were female (55.7%). 74.9% of patients were White (Table 1).
- Majority of specialty of the prescribing provider was primary care physician (53.7%), followed by nurse practitioners (18.4%).
- The majority of GLP-1 RA patients had T2D (87.8%) and 45.5% were obese/overweight.
- Semaglutide (Ozempic, Rybelsus, Wegovy) increased yearly in the Medicare population and accounted for half the GLP-1 RAs initiated during the study period, with Ozempic and Rybelsus accounting for 53.8% and 11.4%, respectively, on or after 2022 (Figure 3). Wegovy was initiated predominantly from March 2024 onwards.
- The 3-year cumulative incidence of myocardial infarction (MI) was 9.5% (95% CI: 9.4–9.6%) (Figure 4a), with numerically varied by drug (Figure 4b).
- Overall survival at 3 years was 87.5% (95% CI: 87.4-87.6%) among GLP-1 RA initiators (Figure 5a), and numerically varied by generic drug (Figure 5b).

## Limitations

- The study was descriptive and does not establish causal relationships between GLP-1 RA use, indications, and death and cardiovascular outcomes.
- The study population was limited to Medicare FFS beneficiaries and may not be representative of other populations (e.g., younger patients, Medicare Advantage).
- Cardiovascular outcomes were identified using administrative claims data, which may be subject to coding inaccuracies.
- Key variables such as BMI, lab values, or lifestyle factors were not available or had high missingness in claims.

## Why is this Research Important

- GLP-1 RAs are medications used to treat Type 2 diabetes and help reduce heart and metabolic risks.
- Semaglutide use increased rapidly over the study period, becoming the most commonly used GLP-1 RA by 2023.
- While endocrinologists can prescribe GLP-1 medications, most patients received theirs from primary care doctors or nurse practitioners.
- This study tracked MI and overall survival over three years among Medicare patients taking GLP-1 medications, providing real-world benchmarks across all GLP-1 indications.
- In this analysis, semaglutide and tirzepatide demonstrated a numerically greater reduction in death and MI compared to other GLP-1 receptor agonists. Further investigation is warranted to understand the factors contributing to these differences.

## Results

Figure 1. Attrition Diagram for Patients Initiating GLP-1 RAs in 100% Medicare FFS Population

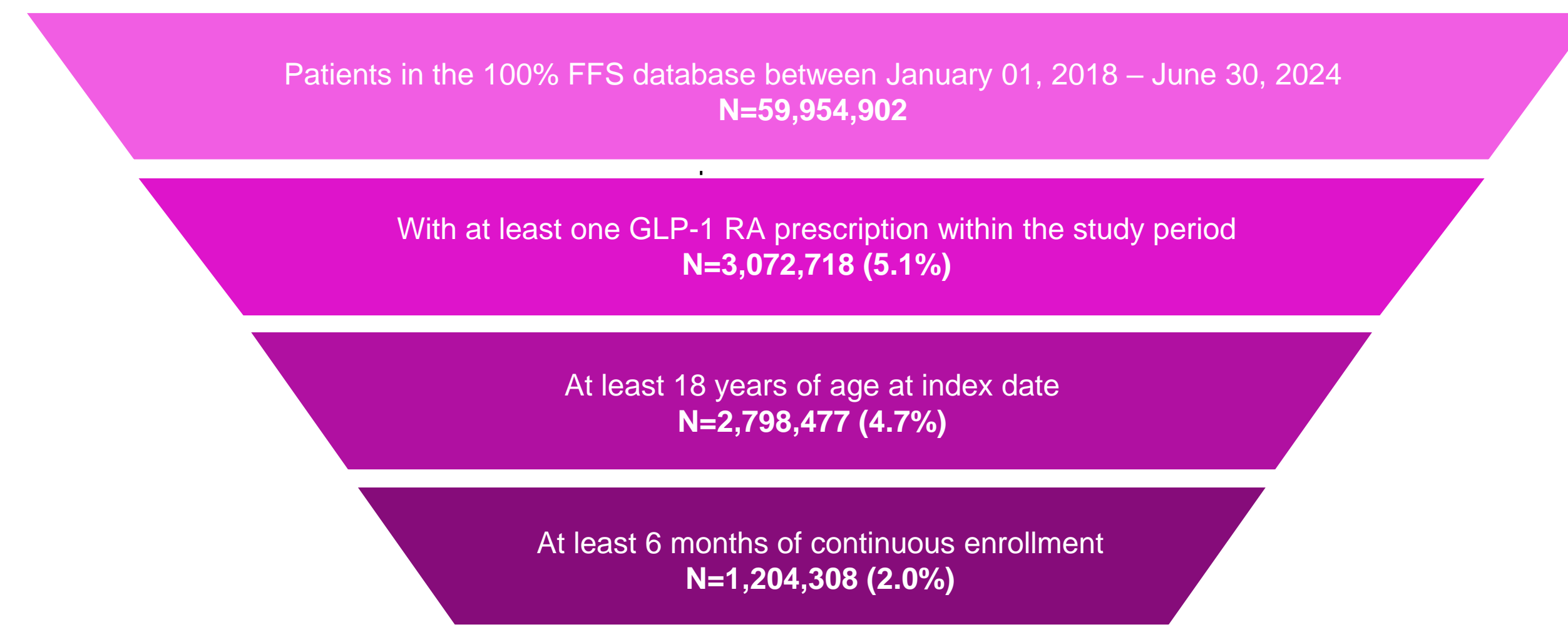
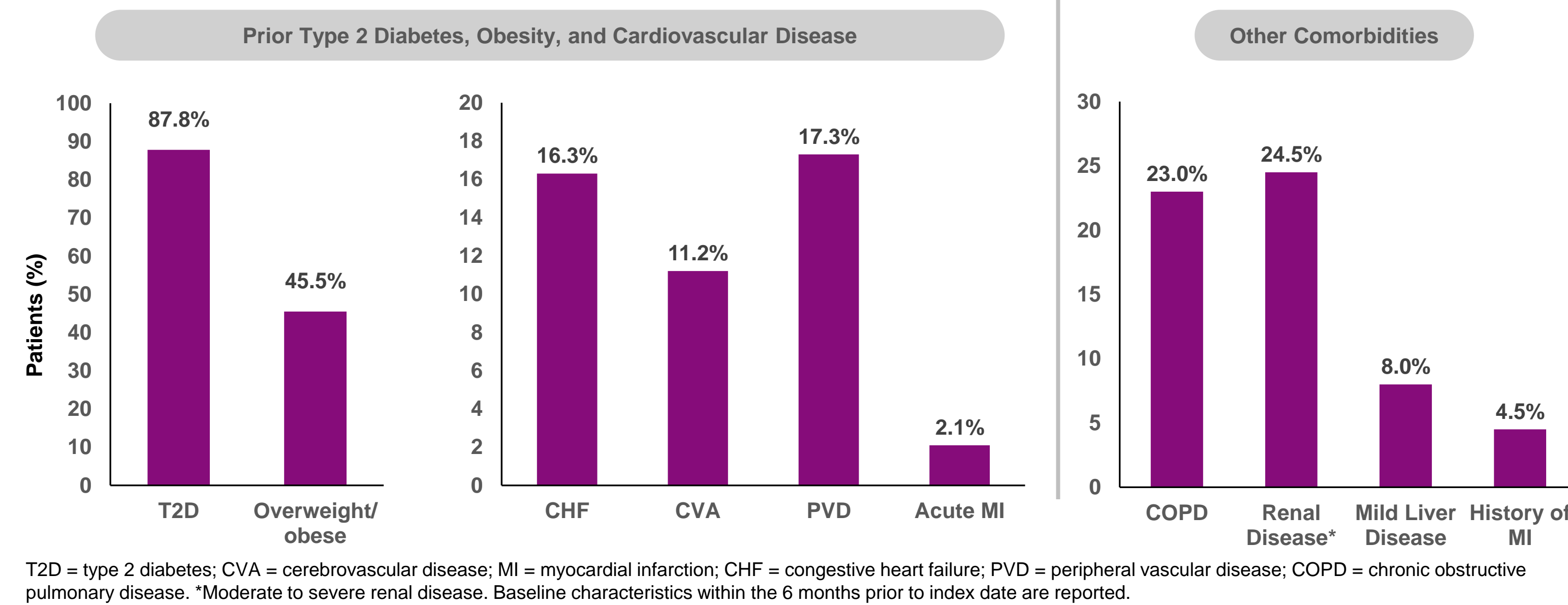


Table 1. Baseline Demographic and Clinical Characteristics

|   | All GLP-1 RA initiators<br>N=1,204,308<br>n (%) |
|---|---|
| <b>Age at GLP-1 RA initiation (years)</b> |   |
| Mean (STD)                                | 67.8 (10.46)                                    |
| Median (Q1-Q3)                            | 69 (65-74)                                      |
| <b>Sex</b>                                |   |
| Male                                      | 533,758 (44.32%)                                |
| Female                                    | 670,549 (55.68%)                                |
| Unknown                                   | 1 (<0.01%)                                      |
| <b>Race/ethnicity</b>                     |   |
| White                                     | 901,533 (74.86%)                                |
| Black                                     | 118,217 (9.82%)                                 |
| Asian                                     | 34,508 (2.87%)                                  |
| Hispanic                                  | 102,621 (8.52%)                                 |
| Other/Unknown                             | 47,429 (3.94%)                                  |
| <b>Region</b>                             |   |
| Midwest                                   | 267,167 (22.18%)                                |
| Northeast                                 | 225,829 (18.75%)                                |
| South                                     | 475,147 (39.45%)                                |
| West                                      | 236,165 (19.61%)                                |
| <b>Specialty</b>                          |   |
| Primary care physician                    | 646,758 (53.70%)                                |
| Nurse practitioner                        | 221,782 (18.42%)                                |
| Endocrinologist                           | 181,264 (15.05%)                                |
| Other                                     | 153,270 (12.73%)                                |
| Unknown                                   | 1,234 (0.10%)                                   |

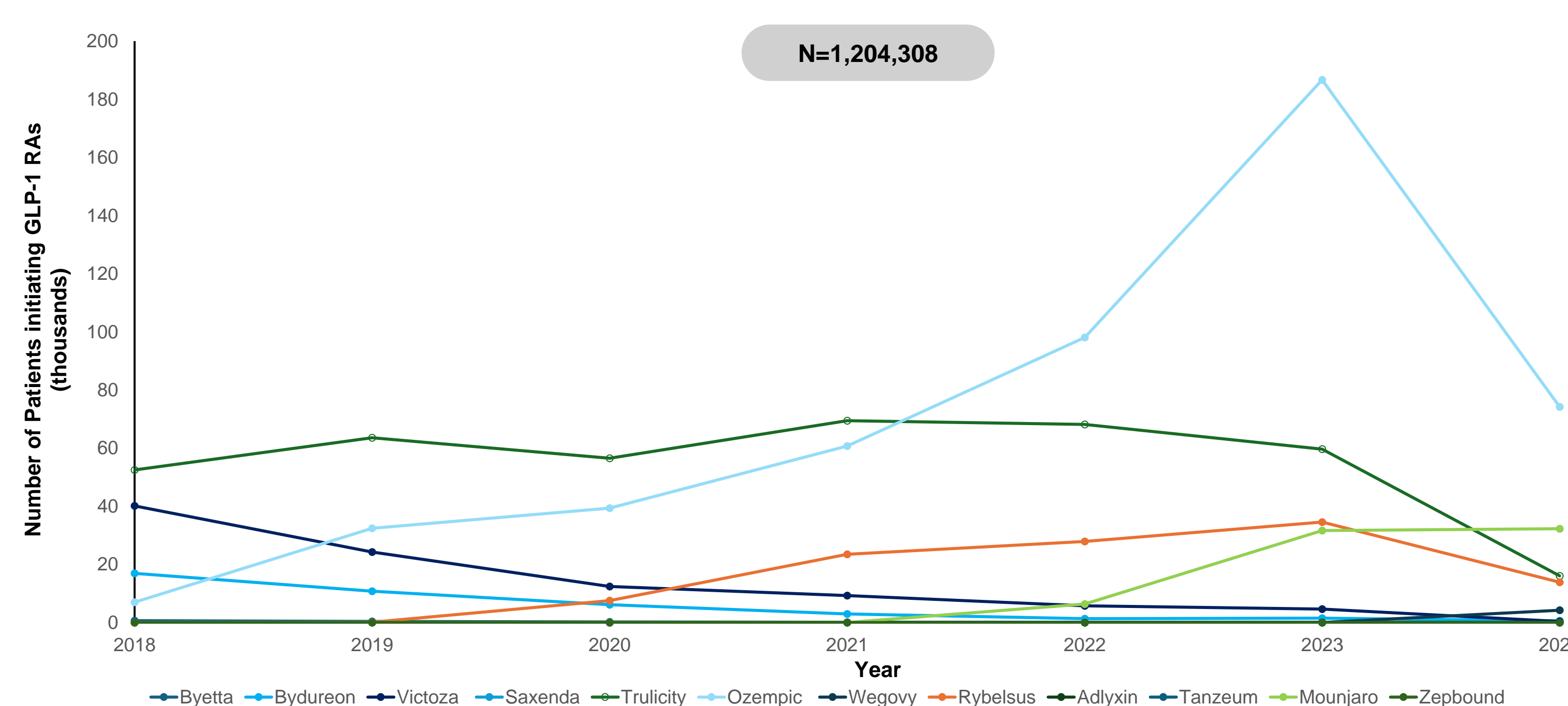
Comorbidities are identified within the 6-month baseline period. Overweight or obesity is derived using ICD-10-CM diagnosis codes. Specialty of the prescribing provider for the index drug is reported. STD = standard deviation; Q1 = quartile 1; Q3 = quartile 3.

Figure 2. Baseline Comorbidities



T2D = type 2 diabetes; CVA = cerebrovascular disease; MI = myocardial infarction; CHF = congestive heart failure; PVD = peripheral vascular disease; COPD = chronic obstructive pulmonary disease. \*Moderate to severe renal disease. Baseline characteristics within the 6 months prior to index date are reported.

Figure 3. Yearly Uptake of GLP-1 RAs in 100% Medicare FFS Population



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### References

References are available upon request to the corresponding author: Shivani@landmarkscience.com

Figure 4. Cumulative Incidence of Myocardial Infarction among GLP-1 RA Initiators

|             | Events/<br>Total | Number<br>Censored | Total PT at<br>risk (years) | 1 year Cumulative<br>Incidence (95% CI) | 2 year Cumulative<br>Incidence (95% CI) | 3 year Cumulative<br>Incidence (95% CI) |
|-------------|------------------|--------------------|-----------------------------|---|---|---|
| Overall     | 60,101/1,062,624 | 1,002,523          | 1,722,938                   | 3.6% (3.6-3.7%)                         | 6.5% (6.5-6.6%)                         | 9.5% (9.4-9.6%)                         |
| Exenatide   | 3,700/40,897     | 37,197             | 115,095.6                   | 3.5% (3.3-3.7%)                         | 6.2% (5.9-6.4%)                         | 8.8% (8.5-9.2%)                         |
| Liraglutide | 9,167/96,383     | 87,216             | 245,099.8                   | 4.1% (4.0-4.2%)                         | 7.4% (7.2-7.6%)                         | 10.4% (10.2-10.7%)                      |
| Dulaglutide | 27,980/369,792   | 341,812            | 756,963.7                   | 4.1% (4.0-4.2%)                         | 7.2% (7.1-7.3%)                         | 10.1% (10.0-10.2%)                      |
| Semaglutide | 23,055/517,576   | 494,521            | 734,180.8                   | 3.2% (3.2-3.3%)                         | 5.8% (5.7-5.9%)                         | 8.6% (8.4-8.7%)                         |
| Tirzepatide | 995/37,976       | 36,981             | 36,475.66                   | 2.7% (2.6-2.9%)                         | 4.7% (3.9-5.5%)                         | NE (NE-NE)                              |

Figure 4a,  
Overall

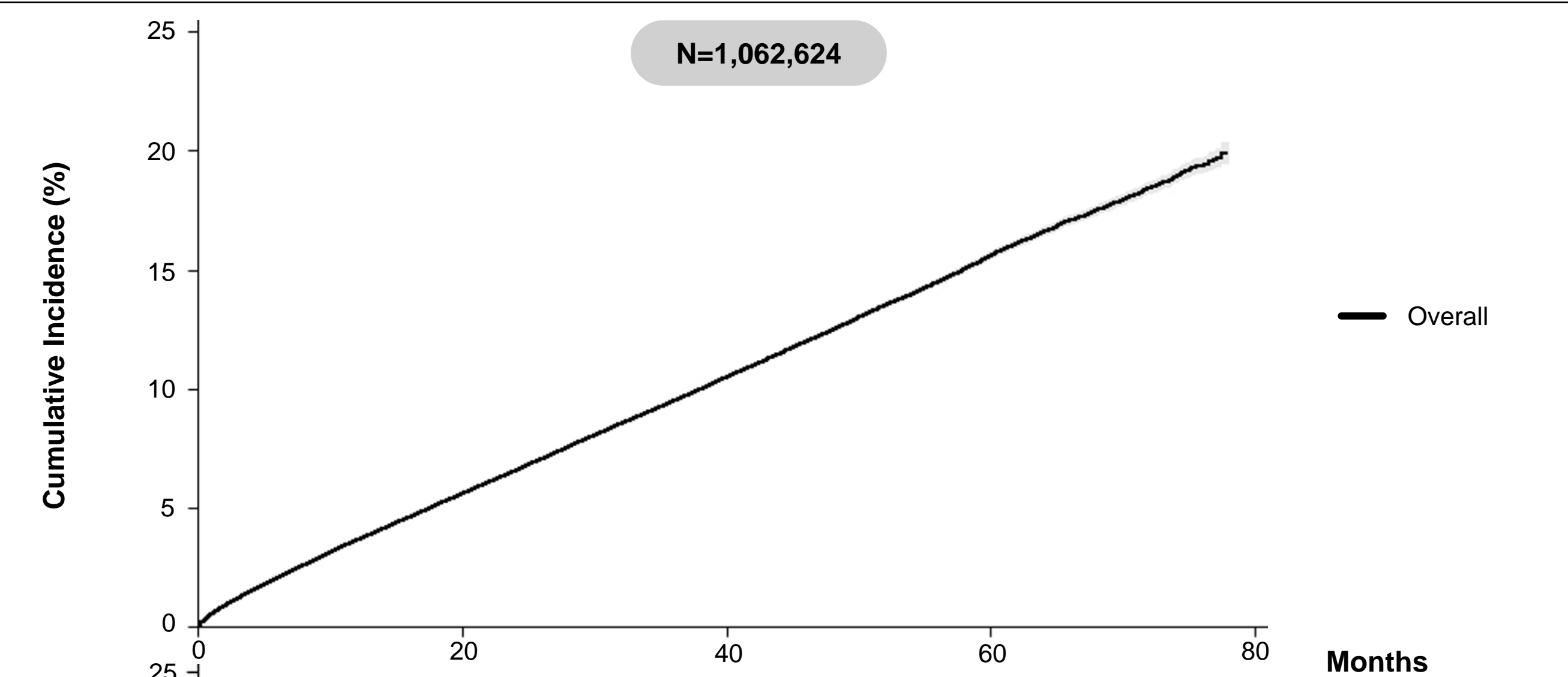
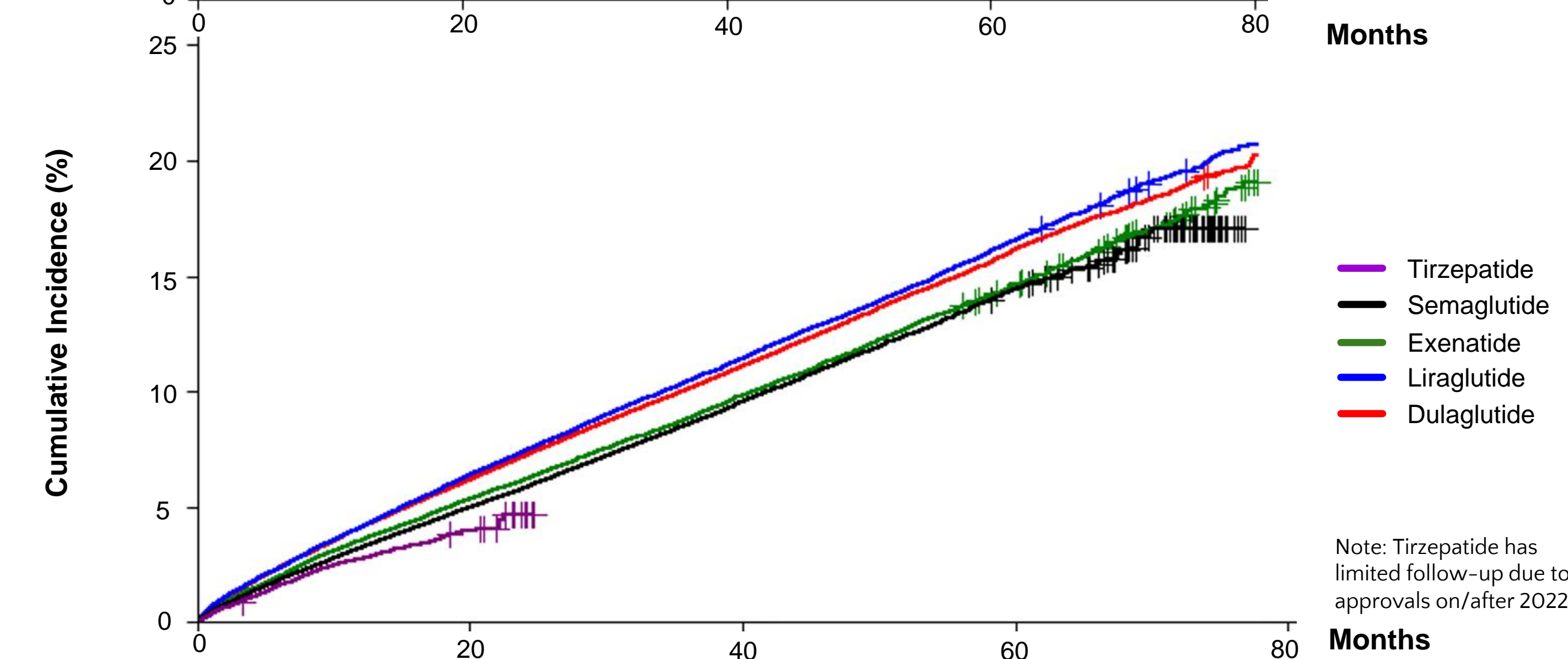


Figure 4b,  
by Generic  
GLP-1 RA



Note: Tirzepatide has limited follow-up due to approvals on/after 2022.

Figure 5. Real-World Overall Survival among GLP-1 RA Initiators

|             | Event/Total      | Number<br>Censored | Total PT at<br>risk (years) | 1 year<br>rwOS (95% CI) | 2 year<br>rwOS (95% CI) | 3 year<br>rwOS (95% CI) |
|-------------|------------------|--------------------|-----------------------------|-------------------------|-------------------------|-------------------------|
| Overall     | 74,355/1,062,624 | 988,269            | 1,791,066                   | 96.9% (96.9-97.0%)      | 92.5% (92.4-92.6%)      | 87.5% (87.4-87.6%)      |
| Exenatide   | 4,777/40,897     | 36,120             | 121,052.8                   | 96.9% (96.8-97.1%)      | 92.7% (92.4-93.0%)      | 88.2% (87.8-88.6%)      |
| Liraglutide | 10,717/96,383    | 85,666             | 260,305.3                   | 96.7% (96.5-96.8%)      | 92.4% (92.2-92.5%)      | 87.5% (87.3-87.8%)      |
| Dulaglutide | 36,486/369,792   | 333,306            | 792,638.5                   | 96.1% (96.0-96.2%)      | 91.4% (91.3-91.5%)      | 86.6% (86.5-86.8%)      |
| Semaglutide | 21,815/517,576   | 495,761            | 756,512.6                   | 97.7% (97.7-97.8%)      | 94.3% (94.2-94.4%)      | 90.3% (90.1-90.4%)      |
| Tirzepatide | 560/37,976       | 37,416             | 37,009.51                   | 98.5% (98.3-98.6%)      | 96.6% (95.9-97.2%)      | NE (NE-NE)              |

Figure 5a,  
Overall

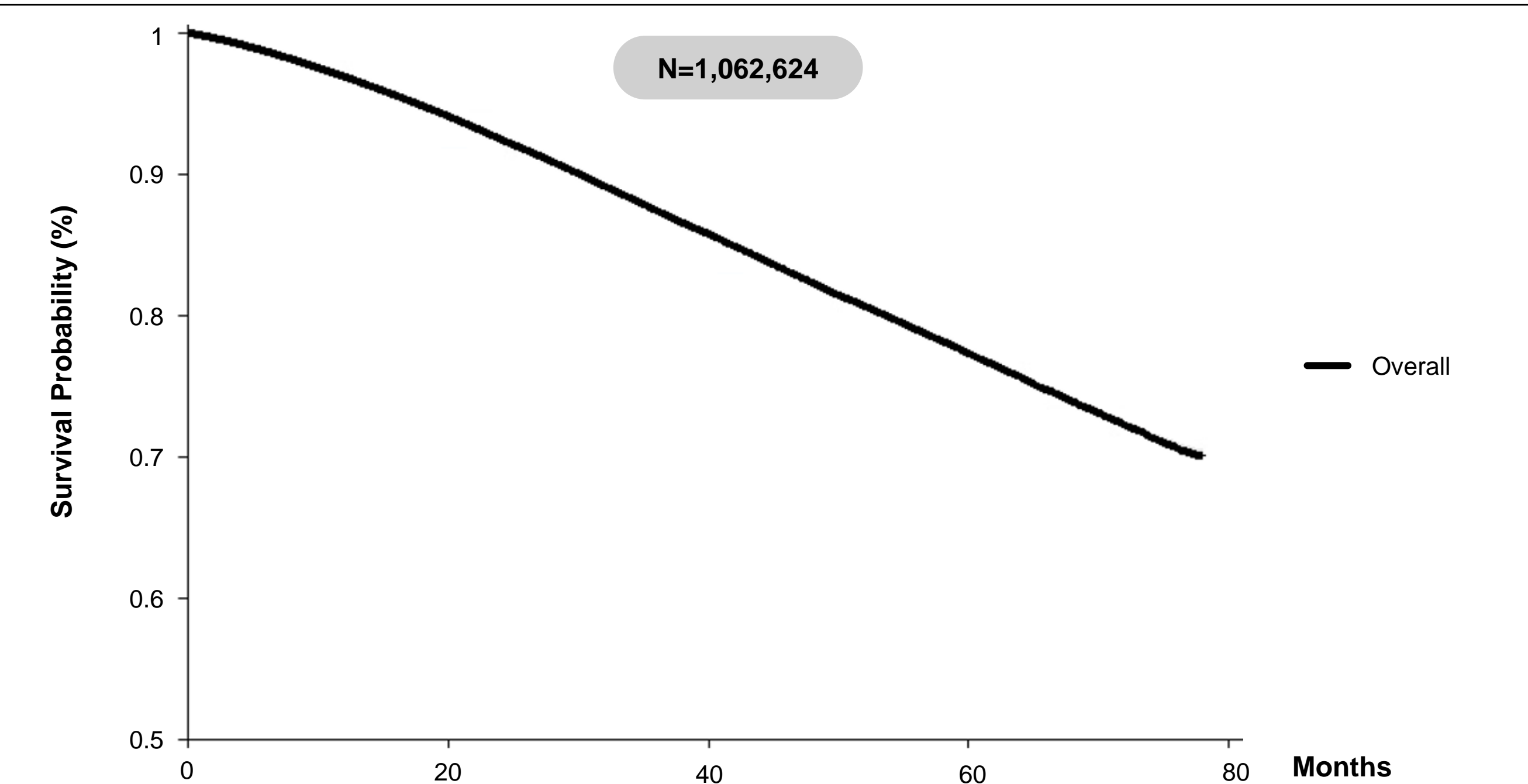
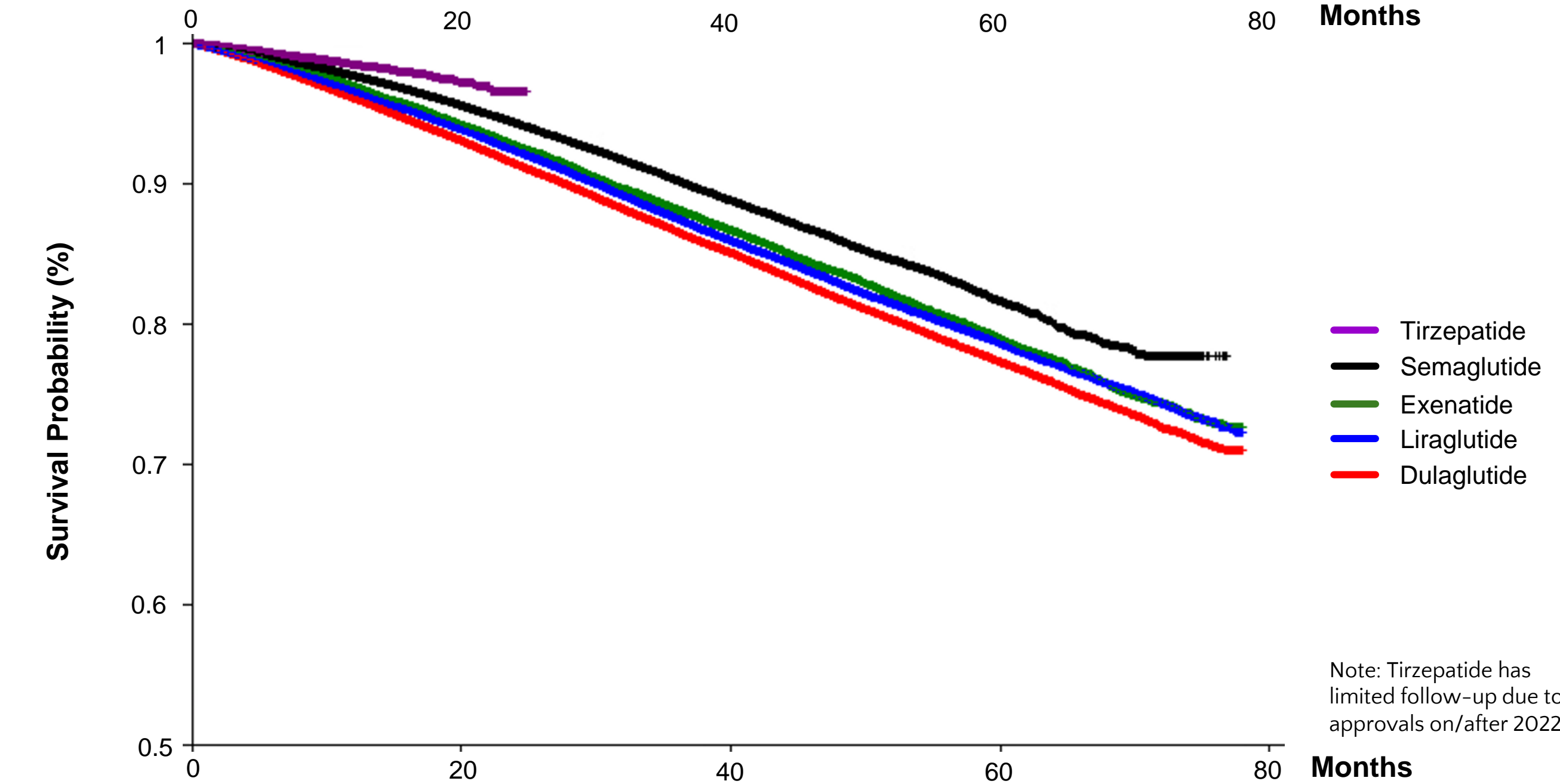


Figure 5b,  
by Generic  
GLP-1 RA



Note: Tirzepatide has limited follow-up due to approvals on/after 2022.

### Funding

This study was funded by Landmark Science, Inc. and Humbi, LLC.

### Disclosures

SA, JW, and DG are employees of or are contracted to Landmark Science, Inc. SV, NA, and PB are employees of Humbi, LLC.