

# USE OF GLUCOSE-LOWERING MEDICATIONS IN THE MANAGEMENT OF TYPE 2 DIABETES



HEALTHY LIFESTYLE BEHAVIORS; DIABETES SELF-MANAGEMENT EDUCATION AND SUPPORT (DSMES); SOCIAL DETERMINANTS OF HEALTH (SDOH)

Goal: Cardiovascular Risk Reduction in High-Risk Patients with Type 2 Diabetes (in addition to comprehensive CV risk management)\*

Goal: Achievement and Maintenance of Glycemic and Weight Management Goals

• AACEVD†

Defined differently across CVDs but all included individuals with established CVD (e.g., MI, stroke, any revascularization procedure). Variably included conditions such as transient ischemic attack, unstable angina, angina pectoris, symptomatic or asymptomatic coronary artery disease.

• Indicators of high risk

While definitions vary, most comprise ≥55 years of age with two or more additional risk factors (including obesity, hypertension, smoking, dyslipidemia, or albuminuria).

• eGFR

Current or prior symptoms of HF with documented BNP or HFrEF

• eGFR

• eGFR ≥30 mL/min per 1.73 m² or SB albuminuria (A2) ≥3.0 mg/annual (30mg/g). These measurements may vary over time, thus, a repeat measure is required to document CKD.

• eGFR

• eGFR with proven HF benefit in this population

• eGFR for maximally tolerated dose of ACEi/ARB

**PREFERABLY**

• eGFR with primary evidence of reducing CVD progression

Use SGLT2i in people with an eGFR ≥20 mL/min per 1.73 m², once initiated should be continued until initiation of dialysis or transplantation

• eGFR

• eGFR with proven CVD benefit if SGLT2i not tolerated or contraindicated

• eGFR

• eGFR, above target, for patients on SGLT2i, consider incorporating a SGLT2i or vice versa

• eGFR with proven CVD benefit

• eGFR with proven CVD benefit

If HbA<sub>1c</sub> above target

• For patients on a SGLT2i RA consider adding SGLT2i with proven CVD benefit or vice versa

• TZD\*

If additional cardiovascular risk reduction or glycemic lowering needed

**Glycemic Management:** Choose approaches that provide the efficacy to achieve goals.

Metformin (or Agent(s) including COMBINATION therapy that provide adequate EFFICACY to achieve and maintain treatment goals. Consider avoidance of hypoglycemia a priority in high-risk individuals.

In general, higher efficacy approaches have greater likelihood of achieving glycemic goals.

Efficacy for glucose lowering

Very High: Dulaglutide (high dose), Semaglutide, Tirzepatide

Insulin

Combination (oral, Combination Injectable (GLP-1 RA/Insulin))

High: GLP-1 RA (not listed above), Metformin, SGLT2i, Sulfonylurea, TZD

Intermediate: DPP-4i

**Achievement and Maintenance of Weight Management Goals:**

Set individualized weight management goals

General lifestyle advice: medical nutrition therapy/behavioral pattern/physical activity

Intensive evidence-based structured weight management program

Consider medication for weight loss

Consider metabolic surgery

When choosing glucose-lowering therapies: Consider regimens with high-to-very-high dual glucose and weight efficacy

Efficacy for weight loss

Very High: Semaglutide, Tirzepatide

High: Dulaglutide, Liraglutide

Intermediate: GLP-1 RA (not listed above), SGLT2i

Neutral: DPP-4i, Metformin

If HbA<sub>1c</sub> above target

Identify barriers to goals:

- Consider DSMES referral to support self-efficacy in achievement of goals
- Consider technology (e.g., diagnostic CGM) to identify therapeutic gaps and tailor therapy
- Identify and address SDOH that impact achievement of goals

\* In people with HF, CVD, established CVD or multiple risk factors for CVD, the decision to use a SGLT2i or SGLT2i with proven benefit should be independent of background use of metformin. A strong recommendation is warranted for people with CVD and a weaker recommendation for those with indicators of high CV risk. Moreover, a higher absolute risk reduction and thus lower number needed to treat are seen at higher levels of baseline risk and should be factored into the shared decision-making process. See text for details. † Low-dose TZD may be better tolerated and similarly effective. ‡ For SGLT2i, CV renal outcomes trials demonstrate their efficacy in reducing the risk of composite MACE, CV death, all-cause mortality, HF, and renal outcomes in individuals with T2D with established high risk of CVD. § For GLP-1 RA, CVDs demonstrate their efficacy in reducing composite MACE, CV death, all-cause mortality, MI, stroke, and renal outcomes in individuals with T2D with established high risk of CVD.

# Physician Guide To Diabetic Treatment

**Michal Rosen-Zvi**



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## **Table of Contents Physician Guide To Diabetic Treatment**

1. Understanding the eBook Physician Guide To Diabetic Treatment
  - The Rise of Digital Reading Physician Guide To Diabetic Treatment
  - Advantages of eBooks Over Traditional Books
2. Identifying Physician Guide To Diabetic Treatment
  - Exploring Different Genres
  - Considering Fiction vs. Non-Fiction
  - Determining Your Reading Goals
3. Choosing the Right eBook Platform
  - Popular eBook Platforms
  - Features to Look for in an Physician Guide To Diabetic Treatment
  - User-Friendly Interface
4. Exploring eBook Recommendations from Physician Guide To Diabetic Treatment

- Personalized Recommendations
- Physician Guide To Diabetic Treatment User Reviews and Ratings
- Physician Guide To Diabetic Treatment and Bestseller Lists
- 5. Accessing Physician Guide To Diabetic Treatment Free and Paid eBooks
  - Physician Guide To Diabetic Treatment Public Domain eBooks
  - Physician Guide To Diabetic Treatment eBook Subscription Services
  - Physician Guide To Diabetic Treatment Budget-Friendly Options
- 6. Navigating Physician Guide To Diabetic Treatment eBook Formats
  - ePub, PDF, MOBI, and More
  - Physician Guide To Diabetic Treatment Compatibility with Devices
  - Physician Guide To Diabetic Treatment Enhanced eBook Features
- 7. Enhancing Your Reading Experience
  - Adjustable Fonts and Text Sizes of Physician Guide To Diabetic Treatment
  - Highlighting and Note-Taking Physician Guide To Diabetic Treatment
  - Interactive Elements Physician Guide To Diabetic Treatment
- 8. Staying Engaged with Physician Guide To Diabetic Treatment
  - Joining Online Reading Communities
  - Participating in Virtual Book Clubs
  - Following Authors and Publishers Physician Guide To Diabetic Treatment
- 9. Balancing eBooks and Physical Books Physician Guide To Diabetic Treatment
  - Benefits of a Digital Library
  - Creating a Diverse Reading Collection Physician Guide To Diabetic Treatment
- 10. Overcoming Reading Challenges
  - Dealing with Digital Eye Strain
  - Minimizing Distractions
  - Managing Screen Time
- 11. Cultivating a Reading Routine Physician Guide To Diabetic Treatment
  - Setting Reading Goals Physician Guide To Diabetic Treatment
  - Carving Out Dedicated Reading Time
- 12. Sourcing Reliable Information of Physician Guide To Diabetic Treatment

- Fact-Checking eBook Content of Physician Guide To Diabetic Treatment
- Distinguishing Credible Sources

### 13. Promoting Lifelong Learning

- Utilizing eBooks for Skill Development
- Exploring Educational eBooks

### 14. Embracing eBook Trends

- Integration of Multimedia Elements
- Interactive and Gamified eBooks

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