

FIRST-LINE Therapy is Metformin and Comprehensive Lifestyle (including weight management and physical activity)



INDICATORS OF HIGH-RISK OR ESTABLISHED ASCVD, CKD, OR HF\*

CONSIDER INDEPENDENTLY OF BASELINE A1C, INDIVIDUALIZED A1C TARGET, OR METFORMIN USE\*

**+ASCVD Indicators of High Risk**

- Established ASCVD
- Indicators of high ASCVD risk (age  $\geq 65$  years with coronary, cerebral, or lower extremity artery disease  $>50\%$ , or LVD)

GLP-1 RA with proven CVD benefit<sup>1</sup> **OR** SGLT2 with proven CVD benefit<sup>2</sup>

If A1C above target

If further intensification is required or patient is unable to tolerate GLP-1 RA and/or SGLT2, choose agents demonstrating CV benefit and/or safety:

- For patients on a GLP-1 RA, consider adding SGLT2 with proven CVD benefit and vice versa<sup>3</sup>
- TZD<sup>4</sup>
- DPP-4 if not on GLP-1 RA
- Basal insulin<sup>5</sup>
- SU<sup>6</sup>

**+HF**

Particularly HF<sup>7</sup> (LVEF  $<45\%$ )

SGLT2 with proven benefit in this population<sup>8,9</sup>

**+CKD**

DND and Albuminur<sup>10</sup>

**PREFERABLY**

SGLT2 with primary evidence of reducing CKD progression

**OR**

SGLT2 with evidence of reducing CKD progression in CVT<sup>11</sup>

**OR**

GLP-1 RA with proven CKD benefit<sup>12</sup> if SGLT2 not tolerated or contraindicated

For patients with T2D and CKD (e.g., eGFR  $<60$  mL/min/1.73 m<sup>2</sup>) and those at increased risk of cardiovascular events

GLP-1 RA with proven CVD benefit<sup>1</sup> **OR** SGLT2 with proven CVD benefit<sup>2</sup>

NO

IF A1C ABOVE INDIVIDUALIZED TARGET PROCEED AS BELOW

**COMPELLING NEED TO MINIMIZE HYPOGLYCEMIA**

DPP-4	GLP-1 RA	SGLT2	TZD
If A1C above target	If A1C above target	If A1C above target	If A1C above target
SGLT2 <b>OR</b> TZD	SGLT2 <b>OR</b> TZD	GLP-1 RA <b>OR</b> DPP-4 <b>OR</b> TZD	SGLT2 <b>OR</b> DPP-4 <b>OR</b> GLP-1 RA

If A1C above target

Continue with addition of other agents as outlined above

If A1C above target

Consider the addition of SU<sup>6</sup> OR basal insulin:

- Choose later generation SU with lower risk of hypoglycemia
- Consider basal insulin with lower risk of hypoglycemia<sup>13</sup>

**COMPELLING NEED TO MINIMIZE WEIGHT GAIN OR PROMOTE WEIGHT LOSS**

GLP-1 RA with good efficacy for weight loss<sup>14</sup> **OR** SGLT2

If A1C above target

If A1C above target

If A1C above target

If quadruple therapy required, or SGLT2 and/or GLP-1 RA not tolerated or contraindicated, use regimen with lowest risk of weight gain

**PREFERABLY**

DPP-4 (if not on GLP-1 RA) based on weight neutrality

If DPP-4 not tolerated or contraindicated or patient already on GLP-1 RA, cautious addition of:

- SU<sup>6</sup> + TZD<sup>4</sup> + Basal insulin

**COST IS A MAJOR ISSUE<sup>15</sup>**

SU<sup>6</sup> **OR** TZD<sup>4</sup>

If A1C above target

TZD<sup>4</sup> **OR** SU<sup>6</sup>

If A1C above target

Health therapy based insulin with lowest acquisition cost

**OR**

Consider other therapies based on cost

1. Proven CVD benefit means it has label indication of reducing CVD events

2. Low dose may be better tolerated though less well studied for CVD effects

3. Dapagliflozin or Glibenclamide have demonstrated CVD safety

4. Choose later generation TZD to lower risk of hypoglycemia; glibenclamide has shown similar CV safety to DPP-4

5. Be aware that SGLT2 labeling varies by region and individual agent with regard to individual level of eGFR for initiation and continued use

6. Dapagliflozin, empagliflozin, and degludec have shown reduction in HF and to reduce CKD progression in CVT. Dapagliflozin and empagliflozin have primary renal outcome data. Dapagliflozin and empagliflozin have others have been shown data

7. Proven benefit means it has label indication of reducing heart failure in this population

8. Refer to Section 11: Musculoskeletal Complications and Foot Care

9. Dapagliflozin / gliclazide (0-100 + gliclazide (0-100) + dulcinate + NPH insulin

10. Sempagliflozin + empagliflozin + dulcinate + aspartate + tolazamide

11. For specific contraindications (e.g., in established CKD), low risk of hypoglycemia, and lower priority to avoid weight gain or no weight-related contraindications

12. Consider country- and region-specific cost of drugs. In some countries TZDs are relatively more expensive and DPP-4 are relatively cheaper

13. Activated whenever these become new clinical considerations regardless of background glucose lowering medications

14. Most patients enrolled in the relevant trials were on metformin or baseline or glucose lowering therapy

# Ada Guidelines Summary

**Brendan G. Carr**



## Ada Guidelines Summary:

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**Background Papers and Summary from the IOM Report**, 1995 [National Summary of State Medicaid Managed Care Programs](#), 2006

**Publications** United States. National Bureau of Standards, 1991 [Current Clinical Medicine E-Book](#) Cleveland Clinic, 2010-08-13 Current Clinical Medicine's 2nd edition by the world famous Cleveland Clinic is an Internal Medicine reference that gives you authoritative and actionable information wherever you are whenever you need it More than 40 updated chapters 13 new chapters and 30% new illustrations ensure that you'll have access to the most up to date guidance In addition to its user friendly easy access format and consistent reliable coverage this Expert Consult title includes a website with the complete contents of the book fully searchable downloadable images and more to keep you and your practice completely current Includes access to a website featuring the complete contents of the book fully searchable access to patient information sheets links to the Gold Standard Drug database and much more to keep you completely current Provides consistent reliable coverage to keep you on the top of your game Includes summary boxes and algorithms for quick confident diagnosis and treatment of each condition Features a user friendly format so you can find information quickly and easily Contains more than a hundred full color illustrations with a special focus on dermatology for highly visual guidance Uses evidence based gradings to help you evaluate your diagnoses Includes many new chapters including Hepatocellular Carcinoma Head and Neck Cancer Takayasu's Arteritis and Non Hodgkin and Hodgkin Lymphoma as well as more than 40 substantially revised chapters that ensure that you'll have access to the most current coverage Features 30% new illustrations that provide you with updated details concepts and procedures

**Americans with Disabilities Act Handbook**, 1991 This guide for architects and builders explains requirements buildings must meet under the Americans with Disabilities Act and includes Final Fair Housing Accessibility guidelines design guidelines for accessible and adaptable dwellings detailed drawings and dimensions for new construction amendments to final ADA guidelines and much more

**Government Reports Annual Index**, 1990 **Federal Register**, 2003-12 **Americans with Disabilities Cases**, 2008 [Americans with Disabilities Act Facilities Compliance](#) Evan Terry Associates, 1993 Helps

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**Labor Relations Reporter** ,2004 Includes sections Decisions of National Labor Relations Board and Court decisions on the law of labor relations Government Reports Announcements & Index ,1992 *Labor Policy and Practice: Labor relations* Bureau of National Affairs (Arlington, Va.),1953 **Computers, Control & Information Theory** ,1986

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