

UPDATED FOR 2025

# Clinical Practice Guidelines Quick Reference Guide



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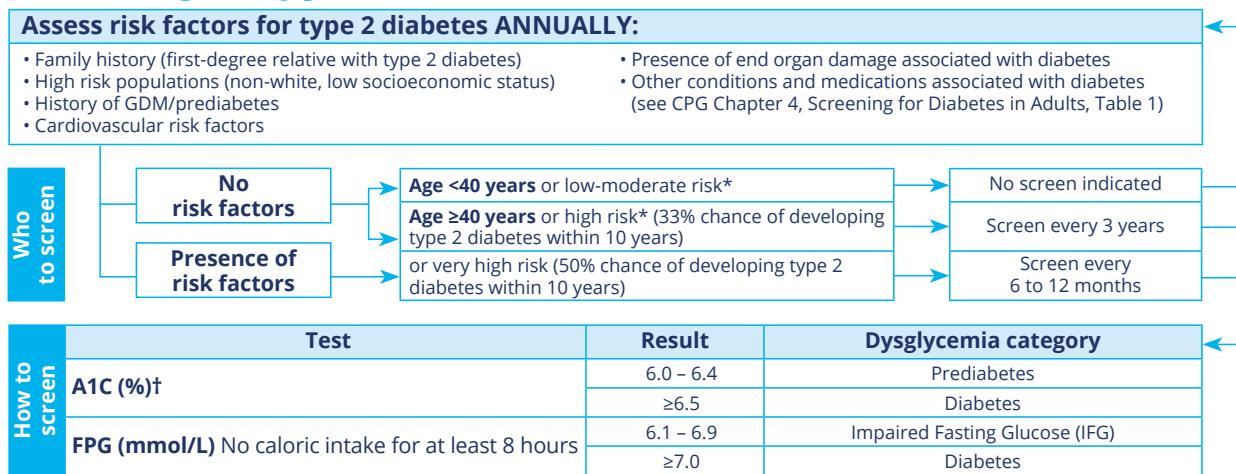
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~~DIABETES~~  
CANADA

## Screening of Type 2 Diabetes

### Assess risk factors for type 2 diabetes ANNUALLY:

- Family history (first-degree relative with type 2 diabetes)
  - High risk populations (non-white, low socioeconomic status)
  - History of GDM/prediabetes
  - Cardiovascular risk factors
- Presence of end organ damage associated with diabetes
- Other conditions and medications associated with diabetes (see CPG Chapter 4, Screening for Diabetes in Adults, Table 1)



## Diagnosis of Diabetes (see CPG "Diabetes and Pregnancy" Chapter for diagnosis of gestational diabetes)

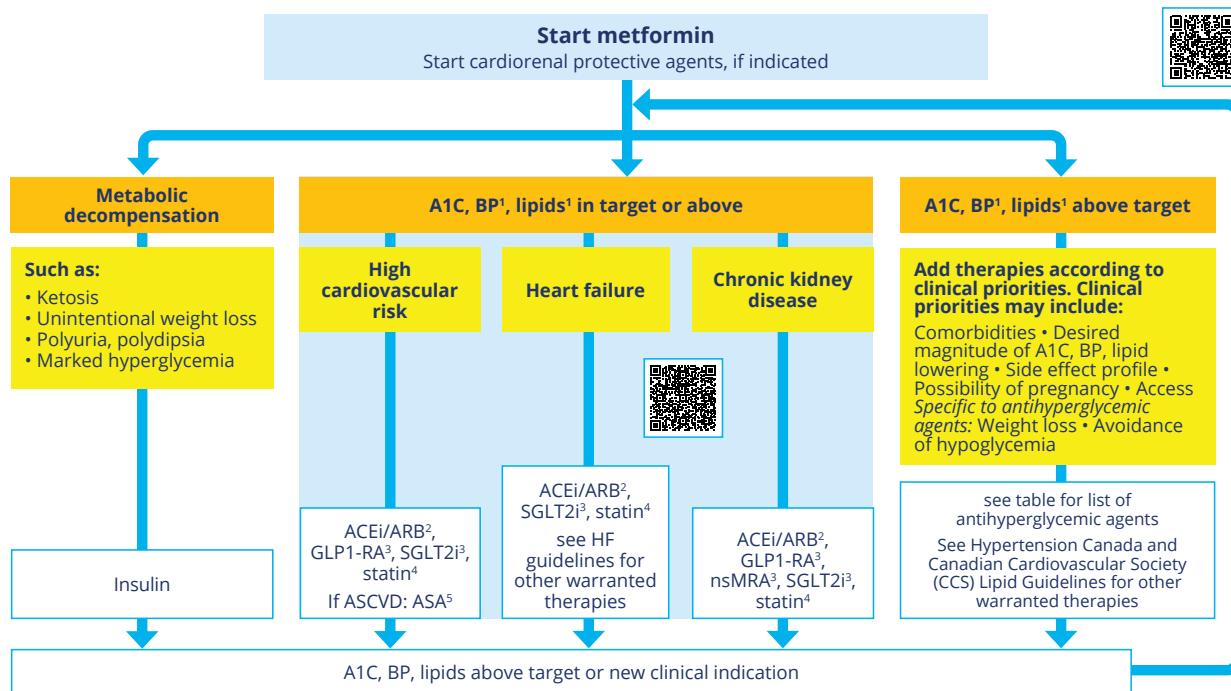
IF	Diagnosis of diabetes	Comments
ASYMPTOMATIC	TWO (2) results (A1C +/- FPG) in the diabetes range – [2 <sup>nd</sup> result confirms the diagnosis in absence of symptoms]	E.g., when one A1C in diabetes range, order a repeat A1C test in a timely manner to confirm the diagnosis of diabetes, or if both A1C and FPG in diabetes range, diagnosis can be made immediately
Symptoms of overt hyperglycemia present <sup>§</sup>	only ONE (1) result in the diabetes range	In addition to A1C and FPG, diagnosis can be made with: 2hPG in a 75g OGTT or Random PG >11.1 mmol/L

\* using a validated risk calculator (e.g., CANRISK)

† Be aware of factors that affect A1C accuracy (see CPG Chapter 9, Table 1)

§ Symptoms of overt hyperglycemia, e.g., polyuria, polydipsia, polyphagia, recent unexplained weight loss

# Pharmacotherapy management for type 2 diabetes



<sup>1</sup> See Hypertension Canada / Canadian Cardiovascular Society (CCS) Lipid Guidelines for other warranted therapies. Treat to BP <130/80 mmHg; lipid targets LDL-C ≤2.0 mmol/L (non-HDL-C ≤ 2.6 mmol/L, apo B ≤ 0.8 g/L); or, with ASCVD, LDL-C ≤1.8 mmol/L (non-HDL-C ≤2.4 mmol/L, apo B ≤0.7 g/L)

<sup>2</sup> ACE-inhibitor or ARB (angiotensin receptor blocker) should be given at doses that have demonstrated vascular protection (e.g., perindopril 8 mg once daily [EUROPA trial], ramipril 10 mg once daily [HOPE trial], telmisartan 80 mg once daily [ONTARGET trial]).

<sup>3</sup> Should be given at doses that have demonstrated vascular protection as tolerated. Not approved by Health Canada for use in type 1 diabetes

<sup>4</sup> See CCS Lipid Guidelines for other warranted therapies.

<sup>5</sup> ASA should not routinely be used for the primary prevention of cardiovascular disease in people with diabetes. ASA may be used for secondary prevention. Consider clopidogrel if ASA-intolerant.

## A1C Targets for glycemic management

A1C (%)	Targets
<6.0	Selected adults with type 2 diabetes with potential for remission to normoglycemia
≤6.5 <sup>1</sup>	Adults with type 2 diabetes to reduce the risk of chronic kidney disease and retinopathy if at low risk of hypoglycemia <sup>2</sup>
≤7.0	<b>MOST PEOPLE WITH TYPE 1 OR TYPE 2 DIABETES</b>
≤8.0	Functionally dependent <sup>3,4</sup>
≤8.5	Frail individuals and/or with cognitive impairment <sup>3,4</sup> Limited life expectancy <sup>3,4</sup>
7.1 – 8.5	Recurrent level 3 hypoglycemia and/or impaired awareness of hypoglycemia
Avoid higher A1C to minimize risk of symptomatic hyperglycemia and acute and chronic complications	

<sup>1</sup> Target A1C 6.0 – 6.4 % for adults with type 2 diabetes with potential for remission to prediabetes

<sup>2</sup> Based on class of antihyperglycemic medication(s) utilized and the person's characteristics

<sup>3</sup> If therapy includes sulfonylurea or insulin, A1C 7.1 % is the recommended lower limit

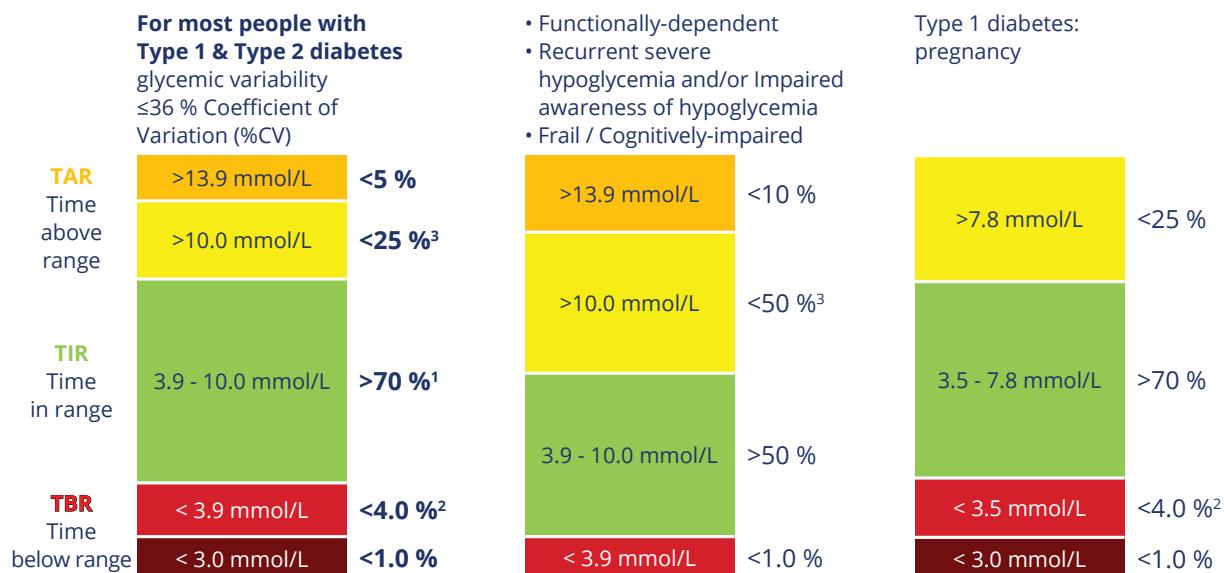
<sup>4</sup> In frail, older adults, especially those living in continuing care homes, may rely less on A1C; focus on avoiding symptomatic hyperglycemia and any hypoglycemia. See Diabetes in Older People chapter.

## Blood Glucose (BG) Targets for glycemic management (when indicated/accessible)

Blood Glucose (BG) Targets	Fasting / Preprandial BG (mmol/L)	2-hr Postprandial BG (mmol/L)
For most people with diabetes If not at A1C ≤7.0 % despite the above BG targets →	4.0 – 7.0 4.0 – 5.5	5.0 – 10.0 5.0 – 8.0

# Continuous Glucose Monitoring (CGM)

## Targets for glycemic management (when indicated\*/accessible)



\* When not at risk of hypoglycemia, may consider targeted, periodic use of CGM in engaged individuals to identify therapeutic gaps, tailor therapy and support individualized daily self-management

<sup>1</sup> Corresponds with an A1C of approximately 7%. Every absolute 10% change in %TIR correlates with 0.5 – 0.8 % change in A1C

<sup>2</sup> includes values smaller than 3.0 mmol/L

<sup>3</sup> includes values greater than 13.9 mmol/L

## Antihyperglycemic Agents and Kidney Function



Maximum Daily Dose of Regular Release Formulation (Unless specified with footnotes)

eGFR (mL/min/1.73 m <sup>2</sup> )	Biguanides		Incretins			SGLT2 Inhibitors			Secretagogues	Others	Insulins		
	Metformin		DPP4 Inhibitors	GIP/GLP1-RA	GLP1-RA	Canagliflozin	Dapagliflozin	Empagliflozin					
Increase frequency of monitoring renal function	≥60	2,550 mg (2,000 mg) <sup>†</sup>	5 mg	5 mg	100 mg	Tirzepatide 15 mg <sup>‡</sup>	Dulaglutide 4.5 mg <sup>‡</sup> Liraglutide 1.8 mg Semaglutide SQ 2 mg <sup>‡</sup> Semaglutide PO 14 mg	300 mg	10 mg	25 mg	Gliclazide 320 mg (120 mg) <sup>§</sup> Glimepiride 8 mg Glyburide 20 mg Repaglinide 12 mg	Acarbose 300 mg Pioglitazone 45 mg	No maximum daily dose
	45-59						100 mg <sup>‡</sup>	No dose change <sup>†</sup>	No dose change <sup>†</sup>	Gliclazide, Glimepiride, Repaglinide - No dose change Avoid glyburide	Dose reduction may be needed		
		1,000 mg	2.5 mg	50 mg					10 mg <sup>‡</sup>	Dose reduction may be needed			
	30-44	500 mg		25 mg	25 mg			Do not initiate but can continue <sup>†</sup>			Pioglitazone - No dose change Acarbose - Limited data available		
								Do not initiate but can continue <sup>†</sup>	10 mg <sup>‡</sup>				
	25-29												
	20-24												
	15-19												
	<15 or Dialysis	Avoid		Avoid		Limited data available	Limited data available			Avoid sulfonylureas			

Yellow = Dose reduction   Red = Avoid   Light blue = Limited data available   Yellow = Do not initiate but can continue

\*Extended release formulation   † Cardiorenal benefits preserved, but reduced glucose-lowering efficacy expected   ‡ Administered weekly

DPP4 = Dipeptidyl peptidase 4; eGFR = estimated glomerular filtration rate; GIP = glucose-dependent insulinotropic polypeptide; GLP1 = glucagon-like peptide-1; RA = receptor agonist; SGLT2 = sodium-glucose cotransporter-2; SQ = subcutaneous; PO = oral.

## Keeping people safe when they are sick or at risk of dehydration



**Re-hydrate** appropriately:  
non-caffeinated, minimal sugar fluids  
- electrolyte replacement solutions (Gastrolyte®, Hydralyte®, Pedialyte®); clear soup; water; diet soda (diet ginger-ale); watered down apple juice

**Hold SADMANS** meds. **Restart** once able to eat/drink normally.

- S** sulfonylureas, other secretagogues
- A** ACE-inhibitors
- D** diuretics, direct renin inhibitors
- M** metformin, MRA\*
- A** angiotensin receptor blockers
- N** non-steroidal anti-inflammatory drugs
- S** SGLT2 inhibitors

\*mineralocorticoid receptor antagonist (finerenone, eplerenone, spironolactone)

## Special considerations regarding pregnancy for people with type 1 or type 2 diabetes

For people planning pregnancy, the following steps taken prior to conception:

- **A1C** 7% or less, but strive for ≤6.5% (ensure contraception until at personalized target)
- **Stop:**
  - Non-insulin antihyperglycemic agents (except metformin and/or glyburide)
  - Statins
  - ACEi/ARB prior to pregnancy, but if overt nephropathy exists, continue until detection of pregnancy
- **Start:**
  - Folic acid 1 mg per day x 3 months prior to conception
  - Insulin if target A1C is not achieved on metformin and/or glyburide (type 2)
  - Other antihypertensive agents safe for pregnancy (Labetalol, nifedipine XL) if hypertension control needed
- **Screen for complications:**
  - Eye appointment, serum creatinine, urine ACR, blood pressure
- Aim for **healthy BMI**
- Ensure appropriate **vaccinations** have occurred
- **Refer** to diabetes clinic

## Common Antihyperglycemic Agents for use in type 2 diabetes

Medication	A1C lowering	Weight	Hypo-glycemia	Other adverse effects	Other therapeutic considerations
<b>1ST LINE FOR MOST PEOPLE</b>					
<b>BIGUANIDES:</b> Decrease hepatic glucose production; increase insulin sensitivity and glucose uptake by muscles and tissues; reduce intestinal glucose absorption					
• Metformin 500mg, 850mg • Metformin ER (extended-release) 500mg, 1000mg	↓↓	↔	negligible risk	• GI side effects, such as diarrhea, abdominal pain, nausea, vomiting • Vitamin B12 deficiency	• diarrhea tends to resolve over time and is minimized with starting low dose, slow titration, taking with meals, and using extended-release preparation • Assess vitamin B12 levels periodically or with symptoms or impaired proprioception or peripheral neuropathy

### 2ND LINE (in alphabetical order)

<b>INCRETINS: DPP4i (physiologic effect):</b> stimulate glucose-dependent insulin secretion, slow gastric inhibiting glucagon release; <b>GLP-1 RA ± GIP RA (physiologic and pharmacologic effect):</b> stimulate glucose-dependent insulin secretion, slow gastric inhibiting glucagon release, slow gastric emptying, enhance satiety					
<b>DPP4 inhibitors:</b> • linagliptin 5mg • saxagliptin 2.5mg, 5mg • sitagliptin 25mg, 50mg, 100mg	↓	↔	negligible risk		• avoid using with other incretins • caution with saxagliptin in people with heart failure
<b>GIP/GLP1 receptor agonists:</b> • tirzepatide 2.5mg, 5mg, 7.5mg, 10mg, 12.5mg, 15mg	↓↓↓	↓↓↓	negligible risk	• GI: nausea, vomiting, diarrhea, constipation • worsening retinopathy seen with rapid A1C lowering	• avoid using with other incretins • contraindicated with personal or family history of medullary thyroid cancer or multiple endocrine neoplasia syndrome type 2

<b>GLP1 receptor agonists:</b> <ul style="list-style-type: none"><li>dulaglutide 0.75mg, 1.5mg</li><li>liraglutide 0.6mg, 1.2mg, 1.8mg</li><li>lixisenatide</li><li>semaglutide (oral) 3mg, 7mg, 14mg</li><li>semaglutide (sc) 0.25mg/0.5mg, 1mg, 2mg</li></ul>			negligible risk	<ul style="list-style-type: none"><li>GI: nausea, vomiting, diarrhea, constipation</li><li>worsening retinopathy seen with rapid A1C lowering</li><li>pancreatitis reported in case reports, but not seen in larger studies</li></ul>	<ul style="list-style-type: none"><li>avoid using with other incretins</li><li>contraindicated with personal or family history of medullary thyroid cancer or multiple endocrine neoplasia syndrome type 2</li></ul>
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**INSULINS:** Regulate metabolism of carbohydrates, fat, and protein; promote absorption (storage) of glucose by tissues; reduce hepatic glucose production and secretion

<b>Basal insulins:</b> <ul style="list-style-type: none"><li>degludec U-100, U-200</li><li>glargine U-100, U-300</li><li>icodec</li></ul> <b>Bolus insulins:</b> <ul style="list-style-type: none"><li>aspart</li><li>aspart (faster-acting)</li><li>glulisine</li><li>lispro U-100, U-200</li></ul> <b>Premixed insulins:</b> <ul style="list-style-type: none"><li>biphasic insulin aspart</li><li>lispro/lispro protamine</li></ul>			significant risk		<ul style="list-style-type: none"><li>potentially greatest A1C reduction</li><li>no maximum dose</li></ul>
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**SECRETAGOGUES:** stimulate receptors on pancreatic  $\beta$ -cells to stimulate endogenous insulin secretion

<b>Sulfonylureas:</b> <ul style="list-style-type: none"><li>gliclazide 80mg</li><li>gliclazide MR (modified-release) 30mg, 60mg</li></ul> <b>Meglitinides:</b> <ul style="list-style-type: none"><li>Repaglinide 0.5mg, 1mg, 2mg</li></ul>			minimal / moderate risk		<ul style="list-style-type: none"><li>relatively rapid glucose-lowering response</li><li>meglitinides reduce postprandial hyperglycemia with minimal, if any, reduction in fasting hyperglycemia</li><li>gliclazide preferred over glyburide due to lower risk of hypoglycemia</li><li>repaglinide contraindicated when co-administered with clopidogrel or with gembifrozil</li></ul>
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**SGLT2 INHIBITORS:** Glucose-lowering effect due to reduced glucose reabsorption by the kidney leading to increased urinary glucose excretion; favourable impacts on renal and heart failure outcomes likely related to renal and systemic hemodynamic effects.

<ul style="list-style-type: none"><li>canagliflozin 100mg, 300mg</li><li>dapagliflozin 5mg, 10mg</li><li>empagliflozin 10mg, 25mg</li></ul>	 efficacy declines as eGFR declines		negligible risk	<ul style="list-style-type: none"><li>genital mycotic infections. women are at higher risk</li><li>rare but important risk for euglycemic diabetic ketoacidosis</li><li>increased risk of fractures with canagliflozin</li><li>increased risk of lower extremity amputation with canagliflozin</li><li>no increased risk of urinary tract infection</li></ul>	<ul style="list-style-type: none"><li>to prevent euglycemia DKA, SGLT2 inhibitors should be held or not used if fasting, if consuming a low-carbohydrate diet, if at risk for volume depletion (diarrhea, sepsis), or prior to major surgery</li><li>caution in those at risk of volume depletion (e.g., loop diuretics)</li><li>dapagliflozin not to be used with bladder cancer</li></ul>
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Not listed in this table: acarbose, insulin (basal NPH intermediate-acting; prandial regular U-100, U-500; premixed regular-NPH), sulfonylureas (glimepiride, glyburide), thiazolidinediones

# Stepwise Approach to Insulin Regimens for People with Type 2 Diabetes



Educate on prevention and management of **hypoglycemia**.  
If cardiorenal comorbidities change, reassess other antihyperglycemic agents.

## DECISION TO INITIATE INSULIN

If glycemic targets not met, or symptomatic hyperglycemia/metabolic decompensation

- Start **basal insulin**
- Titrate basal insulin until fasting glucose in target range
- Continue to optimize other antihyperglycemic agents

Basal



IF GLYCEMIC TARGETS NOT MET

- Add one injection per day of **bolus insulin**, with the largest meal
- Titrate bolus insulin until postprandial glucose in target range
- Consider stopping secretagogues to prevent hypoglycemia

Basal-Plus



IF GLYCEMIC TARGETS NOT MET

- Advance to **multiple injections** of bolus insulin at all meals
- Titrate bolus insulin until postprandial glucose in target range
- Avoid secretagogues

Basal-Bolus or Multiple Daily Injections



## Hypoglycemia: Identifying and Treating

For people using glyburide, gliclazide, repaglinide or insulin



### Signs of hypoglycemia

### Classification of hypoglycemia

### Treatment\*

**Adrenergic (autonomic)**

- Trembling
- Palpitations
- Sweating
- Anxiety
- Hunger
- Nausea
- Tingling

**Level 1**

- Glucose level below normal (often between 3.0 and 3.9 mmol/L)
- Associated with autonomic symptoms
- Without neuroglycopenic symptoms or changes to mental status

**Level 2**

- Glucose level below normal (often <3.0 mmol/L)
- Associated with neuroglycopenic symptoms
- Without significant impact on mental status
- With or without autonomic symptoms

**Level 3**

- Glucose level below normal (regardless of glucose reading)
- Associated with neuroglycopenic symptoms resulting in significantly altered mental/physical status
- Requires assistance to treat

#### Level 1 or 2 hypoglycemia:

Ingest 15 g of carbohydrate, preferably as glucose or sucrose (i.e. tablets or solution). Glucose levels should be retested after 15 minutes and re-treated with another 15 g of carbohydrate if the glucose level remains <3.9 mmol/L

#### Examples of 15 g of carbohydrate:

- 4 x 4 g glucose tablets
- 15 mL (3 teaspoons) or 3 packets of table sugar dissolved in water
- 5 cubes of sugar
- 150 mL juice or regular soft drink
- 6 LifeSavers™
- 15 mL (1 tablespoon) honey

#### Level 3 hypoglycemia:

- Conscious: Treat with oral ingestion of 20 g of carbohydrate, preferably as glucose tablets or equivalent (if capable of swallowing) or 3 mg of glucagon intranasal or glucagon 1 mg SC/IM. Retreat with additional doses after 15 minutes if glucose level remains <3.9 mmol/L
- Unconscious: Treat with glucagon (as above) or 10-25 g (20-50 mL of D50W) of glucose IV. Retreat with additional doses after 15 minutes if glucose level remains <3.9 mmol/L

\* After treatment of hypoglycemia, consume usual meal or snack that is due at that time of the day. If a meal is >1 hour away, consume a snack (including 15 g carbohydrate and a protein source)

# Keeping people with diabetes safe when they are at risk of hypoglycemia

<b>Reduce Driving Risk</b>	<p><b>EDUCATE</b> people at risk of hypoglycemia to drive safely with diabetes</p> <p><b>PREPARE</b> Keep fast-acting sugar within reach and other snacks nearby</p> <p><b>BE AWARE</b> of blood glucose (BG) before driving and every 4 hours during long drives. If BG is below 4 mmol/L, treat</p> <p><b>STOP</b> driving and treat if any symptoms appear</p> <p><b>AFTER</b> treating a low, <b>WAIT</b> until BG is above 5 mmol/L to start driving. Note: Brain function may not be fully restored for some time after blood glucose level returns to normal</p> <p>If a person has impaired awareness of hypoglycemia, he/she must check their BG before driving and every 2 hours while driving, or monitor glucoses with a real-time continuous glucose sensor</p>	
	<p><b>Psychoeducational training</b></p> <ul style="list-style-type: none"> <li>Structured diabetes education programs focused on recognizing and reducing frequency of hypoglycemia</li> </ul> <p><b>Choice of pharmacotherapy</b></p> <ul style="list-style-type: none"> <li>Avoid, reduce dose of, or discontinue pharmacotherapies associated with increased risk of hypoglycemia if appropriate</li> <li>Consider long-acting analogues (insulin glargine-100, glargine-300, detemir, or degludec) over NPH insulin</li> <li>Consider second-generation basal insulin analogues (insulin glargine-300 and degludec) over insulin glargine-100 and detemir to reduce the risk of hypoglycemia, including nocturnal hypoglycemia in type 1 and type 2 diabetes</li> </ul> <p><b>Glucose monitoring</b></p> <ul style="list-style-type: none"> <li>Use of continuous glucose monitoring (CGM) and increased frequency of capillary blood glucose (CBG) monitoring to identify episodes of hypoglycemia</li> </ul> <p><b>Surgical (for type 1 diabetes)</b></p> <ul style="list-style-type: none"> <li>Islet cell transplant</li> <li>Pancreas transplant</li> </ul>	

## ABCDESS of diabetes care

		GUIDELINE TARGET (or personalized goal)
<b>A</b>	<b>A1C</b> with other (CGM*, BG*) glycemic targets *when indicated/accessible	A1C $\leq$ 7.0% (or $\leq$ 6.5% to ↓ risk of CKD and retinopathy) If on insulin or insulin secretagogue, assess for hypoglycemia and ensure driving safety A1C 6.0 - $<$ 6.5% for selected adults with type 2 diabetes with potential remission to prediabetes A1C $<$ 6.0 for selected adults with type 2 diabetes with potential remission to normoglycemia
<b>B</b>	<b>BP</b> targets	BP $<$ 130/80 mmHg If on treatment, assess for risk of falls
<b>C</b>	<b>Cholesterol</b> targets	LDL-C $\leq$ 2.0 mmol/L (or $>$ 50 % reduction from baseline); Alternative: non-HDL-C $\leq$ 2.6 mmol/L, apo B $\leq$ 0.8 g/L If ASCVD, LDL $\leq$ 1.8 mmol/L. Alternative: non-HDL-C $\leq$ 2.4 mmol/L, apo B $\leq$ 0.7 g/L
<b>D</b>	<b>Drugs for CV and/or Cardiorenal protection</b>	<ul style="list-style-type: none"> <li>ASCVD: ACEi/ARB<sup>1</sup>, ASA, GLP1-RA<sup>1</sup>, SGLT2i<sup>1</sup>, statin<sup>2</sup></li> <li>CKD: ACEi/ARB<sup>1</sup>, GLP1-RA<sup>1</sup>, nsMRA<sup>1</sup>, SGLT2i<sup>1</sup>, statin<sup>2</sup></li> <li>HF: ACEi/ARB<sup>1</sup>, SGLT2i<sup>1</sup>, statin<sup>2</sup> (see HF guidelines for other warranted therapies)</li> <li>age <math>\geq</math>55 with <math>\geq</math>1 CV risk factor, or diabetes complications: ACEi/ARB<sup>1</sup>, statin<sup>2</sup></li> <li>age <math>\geq</math>40, age <math>\geq</math>30 and diabetes <math>&gt;</math>15 years, or diabetes complications: statin<sup>2</sup></li> </ul>
<b>E</b>	<b>Exercise</b> goals and healthy <b>Eating</b>	<ul style="list-style-type: none"> <li>150 minutes of moderate to vigorous aerobic activity/ week and resistance exercises 2-3 times/week</li> <li>Follow healthy dietary pattern (eg Mediterranean diet, low glycemic index)</li> </ul>
<b>S</b>	<b>Screening</b>	<ul style="list-style-type: none"> <li>Eye (retinopathy): type 1 - annually; type 2 - every 1-2 years</li> <li>Foot: Monofilament/Vibration yearly or more if abnormal</li> <li>Heart: ECG every 3-5 years if age <math>&gt;</math>40 OR diabetes complications</li> <li>Kidney: Test eGFR and ACR yearly, or more if abnormal</li> <li>Liver: Fib-4 every 1 to 2 years</li> <li>Immunizations: ensure up-to-date as per NACI recommendations</li> </ul>
<b>S</b>	<b>Smoking</b> cessation	If smoker: Ask permission to give advice, arrange therapy and provide support
<b>S</b>	<b>Self-management</b> , stress, sleep, other barriers	<ul style="list-style-type: none"> <li>Set personalized goals (see "individualized goal setting" panel)</li> <li>Assess for stress, sleep, mental health and financial or other concerns that might be barriers to goals</li> </ul>

<sup>1</sup> use agents/doses that demonstrated cardiorenal benefits

<sup>2</sup> see Canadian Cardiovascular Society (CCS) Guidelines for other warranted therapies