Medications for Type 2 Diabetes CDE Exam Preparation



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Clinical Practice Guidelines 2018 CDE Competencies 2018

Guidelines.diabetes.ca



Agenda

Medication for Prediabetes

Oral Medication for Glycemic control

Combination medications

Injectable medication for type 2 diabetes

Medications for hypertension

Medications for cholesterol

Sample questions



Prediabetes- Lifestyle

- 58% reduction in progression at 4 years (DPP)
 - Lifestyle- diet, 5% weight reduction, exercise
- 67% reduction in progression Japanese Study
 - Lifestyle for IGT
- 43% reduction Chinese Da Qing Prevention
 - Lifestyle



Medications used for Prediabetes

- Metformin 850 mg bid (DPP) 31% decrease in progression to diabetes
- Acarbose 100 mg tid (STOPNIDDM) 36% reduction
- GLP1 Liraglitide prevalence of prediabetes decreased 84-96% depending on dose
- TZD (ACTNOW)- pioglitazone decreased conversion to diabetes by 72%
- Metformin + rosiglitazone 66% reduction
- Glargine ORIGIN 31% reduction in diabetes



Competency for CDE Exam 3A

Oral Medications for Type 2 Diabetes

- Action
- Indications for Use
- Side Effects
- A1C lowering
- Weight
- Hypoglycemia
- Precautions
- Comments



Changes in 2018 CPG

Cardiovascular Protection

Weight neutral

Prevent Hypoglycemia



There an app for that!



he Canadian Diabetes Association has become Diabetes Canada*



Guidelines

2018 Full Guidelines
Appendices
Quick Reference Guide
Slides
Videos
Get the App

Key Messages

Reduce Complications Keep People Safe Self-management

Diabetes Canada Clinical Practice Guidelines App

The CPGs on Your Smartphone and Tablet

This **FREE** mobile app is designed to incorporate the professional healthcare tools, clinical practice guidelines chapters, slide decks, and narrated slide presentations.

The DC CPG App is available for **iOS** from the App Store and **Android OS** from Google Play. Please click the links below to download the App.







Pharmacologic Glycemic Management of Type 2 Diabetes in Adults

- Nood Glucose-Lowering Therapies (Type 2 Diabetes) Quick Reference
- Examples of Insulin Initiation and Titration Regimens in People with Type 2 Diabetes
- Insulin Guide. What you need to know Video
- Insulin Prescription Tool
- Keeping Patients Safe When They Are At Risk of Dehydration (Vomiting/Diarrhea) Quick Reference
- Keeping Patients Safe Who Are At Not Of Hypoglycemia Quick Reference
- Pharmacotherapy for Type 2 Diabetes Interactive
- Insulin Pen Start Checklist
- Sick Day Medication List
- Therapeutic Considerations for Renal Impairment
- Types of Insulin
- Which Vascular Protection Medication Are Indicated for My Patient Quick Reference
- Ramadan and Diabetes



Is this patient newly diagnosed with type 2	diabetes?
○ Yes ● No	
Are glycemic targets now being met?	
○ Yes ● No	
Does this patient have clinical cardiovascul	ar disease?
● Yes ○ No	
What medications is this patient currently t	aking?
 □ Acarbose ☑ DPP4-i □ GLP1RA □ Insulin - basal □ Insulin - mealtime Submit	 Meglitinide ✓ Metformin SU SGLT2i TZD



Recommendation:

Start antihyperglycemic agent with demonstrated CV benefit (empagliflozin, liraglutide, canagliflozin).

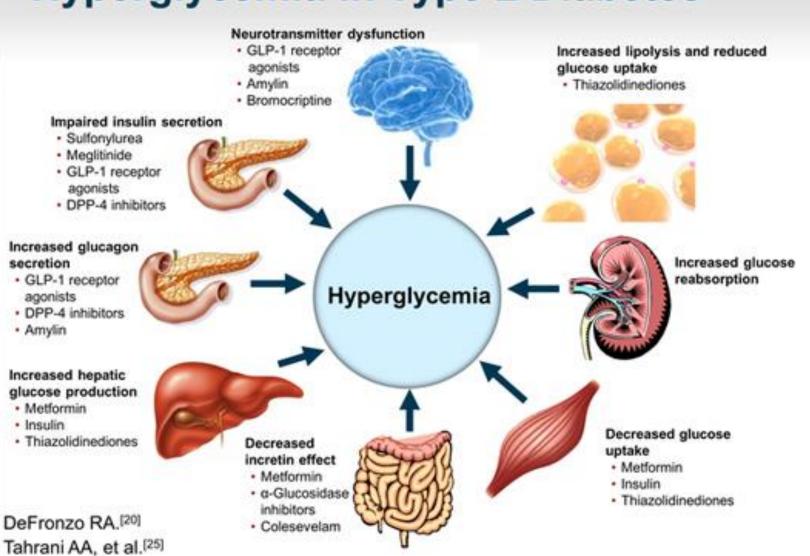


Know this chart!

	Class▲	Effect on CV outcomes (people with diabetes and CVD)	Likelihood of hypoglycemia	Effect on weight	Relative A1C lowering when added to metformin	Cost	Interactions	Renal considerations	Other therapeutic considerations
	Alpha-glucosidase inhibitor (acarbose)		Rare	Neutral	1	\$\$			GI side effects; requires TID dosing
	DPP-4 Inhibitors	Neutral: alogliptin, saxagliptin, sitagliptin	Rane	Neutral	11	\$55			Caution with saxagliptin and heart failure; Rare joint pain
	GLP1 receptor agonists	Liraglutide: Superior Exenatide LAR and lixisenatide: Neutral	Rare	11	11to 111	\$5\$5			GI side-effects Gallstone disease Contraindicated: Personal or family history of MTC or MEN2 Requires SC injection
	Insulin	Neutral: glargine Non-inferior to glargine: degludec	High	**	1111	5-5555			No dose ceiling, flexible regimens Requires SC injection
	Insulin secretagogue: Meglitinide		Some	t	11	\$\$			Reduced post- prandial glycemia; requires TID-QID dosing
	Insulin secretagogue: Sulfonylurea		Some	+	11	\$			Gliclazide and glimepiride associated with less hypoglycemia than glyburide Poor durability
	SGLT2 inhibitors	Canagliflozin and empagliflozin: superior	Rare	11	11:0111	\$5\$			Genital infections, UTI, hypotension Caution with renal dysfunction and loop discretics, elderly Contraindications: Dapagifflozin and bladder cancer, canagifflozin and prior lower extremity amputation Rare DXA (may occur without hyperglycemia)
	Thiazoladinediones	Neutral	Rare	++	11	55			CHF, edema, fractures, cardiovascular controversy (rosiglitazone) Contraindications: pioglitazone and bladder cancer



Hyperglycemia in Type 2 Diabetes





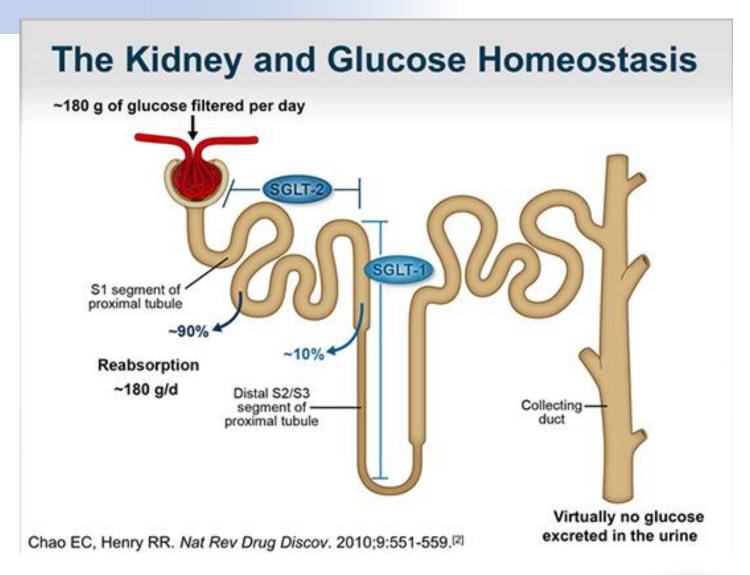
Biguanides

	Class	Drugs	Brand name (non-exhaustive list)	Commercial presentation		Risk of hypoglycemia
0	Biguanides	Metformin	Glucophage	HMR 500 mg	850 mg	No
0	Biguanides	Extended release metformin	Glumetza	M 500 500 mg	M1000 1000 mg	No

Monitor Vitamin B12 levels



SGLT2





SGLT2

Class	Drugs	Brand name (non-exhaustive list)	Comm presen	
Inhibitor of sodium glucose co-transporter 2 (SGLT2)	Canagliflozin	Invokana	100 mg	300 mg
O Inhibitor of sodium glucose co-transporter 2 (SGLT2)	Dapagliflozin	Forxiga	5 mg	10 mg
O Inhibitor of sodium glucose co-transporter 2 (SGLT2)	Empagliflozin	Jardiance	S 10	S 25 25 mg

\$100/ month

Increase fluid intake

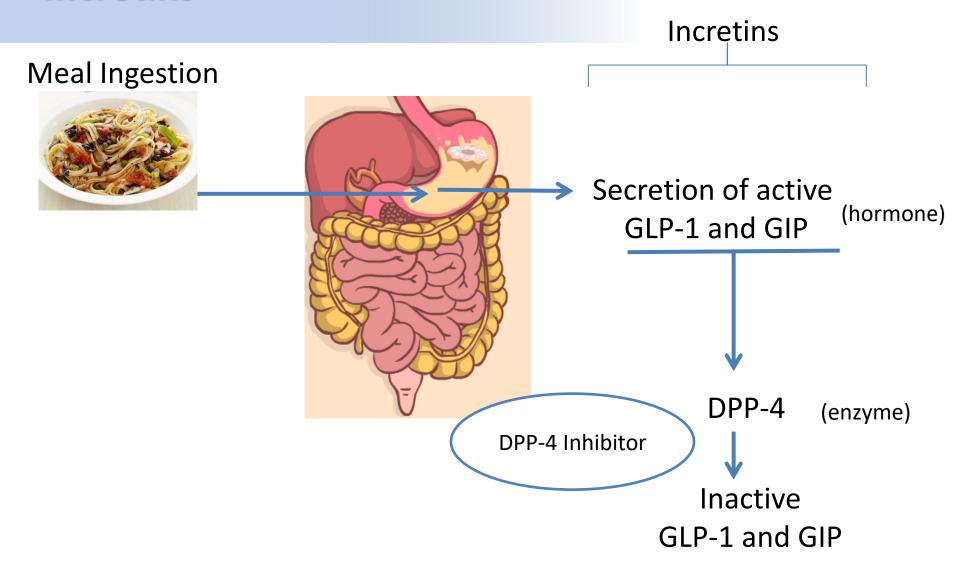


Combinations with SGLT2

Canagliflozin + Metformin	Invokamet	50/500 mg, 50/850mg, 50/1000mg 150/500mg, 150/850 mg, 150/1000mg
Dapagliflozin + Metformin	Xigduo	5/850 mg, 5/1000mg
Empagliflozin + metformin	Synjardy	5/500 mg, 5/850 mg, 5/1000 mg 12.5/500 mg, 12.5/850 mg, 12.5/ 1000mg
Empagliflozin + Linagliptin	Glyxambi	10/5 mg, 25/5 mg



Incretins





DPP-4 Inhibitors

\$100/ month

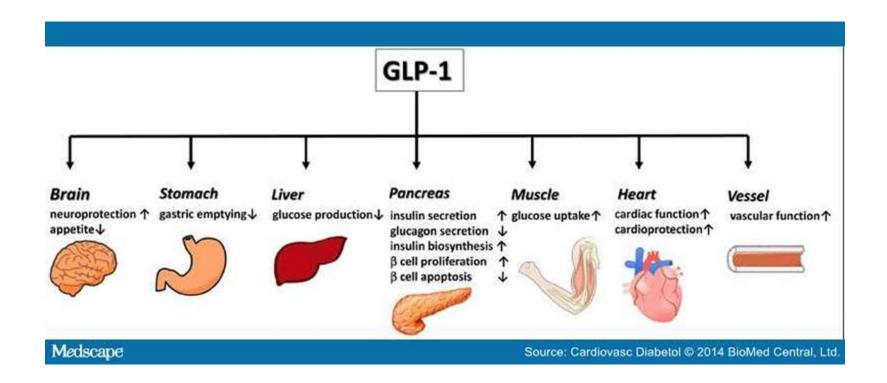
Class	Drugs	Brand name (non-exhaustive list)	Commercial presentation		
O Dipeptidyl peptidase-4 inhibitors (incretin pathway)	Alogliptin	Nesina	6.25 mg 12.5 mg 25 mg		
O Dipeptidyl peptidase-4 inhibitors and biguanides	Alogliptin and metformin	Kazano	12.5/500 mg 12.5/850 mg 12.5/1000 mg		
O Dipeptidyl peptidase–4 inhibitors (incretin pathway)	Linagliptin	Trajenta	5 mg		
O Dipeptidyl peptidase-4 inhibitors and biguanides	Linagliptin and metformin	Jentadueto	2.5/500 mg 2.5/850 mg D2/1000 2.5/1000 mg		
O Dipeptidyl peptidase-4 inhibitors (incretin pathway)	Saxagliptin	Onglyza	2.5 4215		
O Dipeptidyl peptidase-4 inhibitors and biguanides	Saxagliptin and metformin	Komboglyze	2.5/1000 2.5/1000 g 2.5/850 mg 2.5/1000 mg		
O Dipeptidyl peptidase-4 inhibitors (incretin pathway)	Sitagliptin	Januvia	25 mg 50 mg 100 mg		
O Dipeptidyl peptidase-4 inhibitors and biguanides	Sitagliptin and metformin	Janumet	575 50/500 mg 50/850 mg 50/1000 mg		
O Dipeptidyl peptidase-4 inhibitors and biguanides	Extended release sitagliptin and metformin	Janumet XR	50/1000 mg		

Combinations with DPP4

Alogliptin + Metformin	Kazano	12.5/500,mg 12.5/850 mg, 12.5/1000 mg
Linagliptin + Metformin	Jentadueto	2.5/500 mg, 2.5/850 mg, 2.5/1000 mg
Saxagliptin + Metformin	Komboglyze	2.5/500mg, 2.5/850 mg, 2.5/1000 mg
Sitagliptin + metformin	Janumet	50/500 mg, 50/850 mg, 50/1000 mg
Extended release sitagliptin + metformin	Janumet XR	50/500 mg, 50/1000mg, 100/1000mg



GLP-1





GLP-1

Exenatide	Byetta
Liraglutide	Victoza
Lixisenatide	Adlyxine

Not covered by ODB \$168-303/month



GLP-1

Once per week injectable Bydureon (exenatide extended release)



Trulicity (dulaglutide)



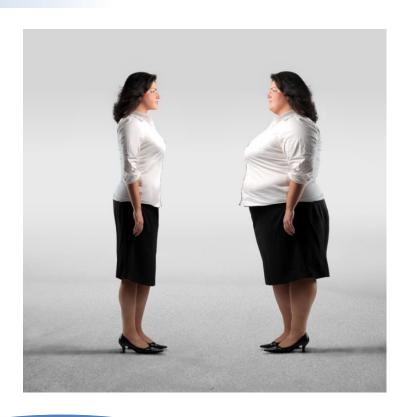




GLP-1 Weight Loss

Saxenda- liraglutide





Not covered by ODB Up to \$413/ month



Secretagogues

Meglitinide

faster acting

Sulfonylurea

longer duration of action

Must be taken with food



Secretagogues

Class	Drugs	Brand name (non-exhaustive list)		mercial entation
Amino acid derivate (insulin secretagogues)	Nateglinide	Starlix	60 mg	120 mg
Meglitinides (insulin secretagogues)	Repaglinide	GlucoNorm	0.5 mg	1 mg 2 mg





Secretagogues

Class	Drugs	Brand name (non-exhaustive list)		nercial ntation
Sulfonylureas (insulin secretagogues))	Glimepiride	Amaryl		2 mg 4 mg
Sulfonylureas (insulin secretagogues)	Glyburide	Diaßeta	2.5 mg	5 mg
Sulfonylureas (insulin secretagogues)	Gliclazide	Diamicron	80 mg	
Sulfonylureas (insulin secretagogues)	Gliclazide modified release	Diamicron MR	DIA 30 30 mg	60 mg

Hypoglycemia Risk





Class	Drugs	Brand name (non-exhaustive list)		mmercial sentation
O Thiazolidinediones	Pioglitazone	Actos	15 mg	30 mg 45 mg
O Thiazolidinediones	Rosiglitazone	Avandia	2 mg	4 mg 8 mg
Thiazolidinediones and biguanides	Rosiglitazone and metformin	Avandamet	2/500 2/500 mg 4/500 4/500 mg	2/1000 2/1000 mg 4/1000 mg



Alpha-glucosidase Inhibitors

Class	Drugs	Brand name (non-exhaustive list)	Comm presen	
Alpha-glucosidase inhibitors	Acarbose	Glucobay	50 mg	100 mg





Competency for CDE Exam 3H, 5F

Cardiovascular Protection

- Medications for Hypertension
- Medications for Cholesterol



Cardiovascular Protection

Prescription for Cardiovascular Protection with diabetes

Prescriber's Name: Address:		Patient's Name: Address:	

STEP 1:	STEP 2: Ch	hoose Cardiovascular protection agent(s) from the following list			Dosing	
Is the patient - age >40? OR - age >30, and diabetes >15 years? OR - warranted for statin therapy based on the Canadian Cardiovascular Society Lipid Guidelines?	Statin	□ Atorvastatin (Lipitor®) □ 10 mg (start 10 mg OD) □ 20 mg □ 40 mg □ 80 mg (max 80 mg OD) □ Pravastatin (Pravachol®) □ 10 mg (start 10 mg OD)	STATIN	☐ 40 mg (max 80 mg OD)	Dosing: see start and maximum doses listed for each statin	
		20 mg 40 mg 80 mg (max 80 mg OD)	□ 10 mg (start 10 mg OD □ 20 mg □ 40 mg (max 40 mg OD	20 mg 40 mg (max 80 mg OD)		
Is the patient	Statin + ACEi or ARB	ACE INHIBITORS		ARB	Dosing: see start and maximum doses listed for each ACEI	
- age >55 with additional CV risk factors? Does the patient have microvascular disease? - Retinopathy - Kidney disease (ACR>2.0) - Neuropathy		□ Perindopril (Aceon®, Coversyl®) □ 2 mg □ 4 mg (start 4 mg OD) □ 8 mg (max 16 mg OD)	□ Ramipril (Altace*) □ 1.25 mg □ 2.5 mg (start 2.5 mg Ol □ 5 mg □ 10 mg (max 20 mg OD	☐ 80 mg (max 80 mg OD)	ACEI: see precautions for dosing in kidney and liver disease on next page Increase doses at 2-3 week intervals.	
Does the patient have cardiovascular disease?	Statin	ASA (if CVD)				
Cardiac disease: Cardiac ischemia (silent or overt) Peripheral arterial disease Cerebrovascular/carotid disease	+	□ ASA □ 81 mg □ 162 mg	☐ Clopidrogrel (Plavix*) for those unable able to tolerate ASA☐ 75 mg			
YES	Statin + ACEi or ARB + ASA + SGLT-2i or GLP-1ra	SGLT-2 inhibitor		Starting dose: lowest dose and titrate		
AND the patient has type 2 diabetes and is NOT at		□ Canagliflozin (Invokana*) □ Empagliflozin (Jardiance in 100 mg (Start 100 mg OD) □ 100 mg (start 100 mg OD) □ 10 mg (Start 10 mg OD) □ 300 mg (max 300 mg OD) □ 25 mg (max 25 mg OD)		(start 10 mg OD)	up Q 4 weeks. Check eGFR periodically, discontinue if eGFR <30mL/min. See benefits and precautions on next page	
glycemic target YES		GLP-1 receptor agonist		Starting dose: Start at 0.6 mg s.c. OD, increase by 0.6 mg Q weekly until		
		□ Liraglutide (Victoza*) □ 0.6 mg (start 0.6 mg OD) □ 1.2 mg □ 1.8 mg (max 1.8 mg OD)			maximum dose reached. If nausea experienced, reduce dose down by mg and use slower titration schedul (Q 2 – 3 weeks between increases) See benefits and precautions on next page	



Know this chart!

Prescription for Cardiovascular Protection with diabetes

STEP 1:	STEP 2: Ch
Is the patient - age >40? OR - age >30, and diabetes >15 years? OR - warranted for statin therapy based on the Canadian Cardiovascular Society Lipid Guidelines?	Statin
Is the patient - age >55 with additional CV risk factors? Does the patient have microvascular disease? - Retinopathy - Kidney disease (ACR>2.0) - Neuropathy	Statin + ACEi or ARB
Does the patient have cardiovascular disease? - Cardiac ischemia (silent or overt) - Peripheral arterial disease - Cerebrovascular/carotid disease	Statin + ACEi or ARB + ASA

Signature:



Is the patient

☐ Age >40

Or

 \square Age >30 + diabetes x 15 years

Or

Warranted for statin based on Canadian
 Cardiovascular Society Lipid Guidelines







Is Patient:

- □ Age >55 with additional CV risk factorsOr
- ☐ Have microvascular disease, retinopathy, kidney, neuropathy





Does the patient have cardiovascular disease

- cardiac ischemia
- peripheral vascular disease
- cerebrovascular /carotid disease





Medications for Hypertension

Target 130/80 mmHg





Medications for Hypertension

Who do you treat?

- Over 55, use an ACE or ARB
- Under 55, with PAD, CVD, microvascular or macrovascular complications, use an ACE or ARB
- Anyone whose Blood pressure is above the target,

use an ACE or ARB





Medications for Hypertension

Combination of 2 first line drugs may be considered as initial therapy if the blood pressure is above target:

≥ 20 mmHg systolic or

≥ 10 mmHg diastolic

Three drugs may be required to reach target.



Medications for Hypertension- Monitoring

Monitor serum potassium and creatinine in patients with CKD prescribed an ACEI or ARB.

Combinations of ACEI and ARB are generally not recommended in the absence of proteinuria.



Medications for Hypertension- ACE (Angiotensin Converting enzymes)

Generic Name	Brand Name
Quinapril	Accupril, generic
Ramipril	Altace, generic
Captopril	Capoten, generic
Perindopril	Coversyl
Benazepril	Lotensin, generic
Cilazapril	Inhibace, generic
Lisinopril	Prinivil, Zestril, generic
Fosinopril	Monopril, generic
Enalapril	Vasotec, generic
Trandolapril	Mavik



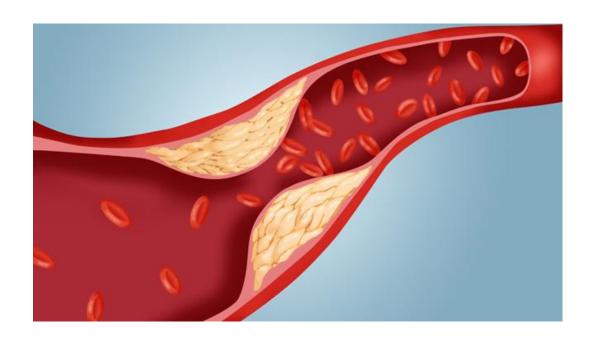
Medications for Hypertension ARB (Angiotensin II Receptor Blockers)

Generic Name	Brand Name
Candesartan	Atacand
Eprosartan	Teveten
Irbesartan	Avapro
Losartan	Cozaar
Telmisartan	Micardis
Valsartan	Diovan
Olmersartan medoxomil	Benicar
Azilsartan	Edarbi



Medications for Hyperlipidemia

Target LDL-C ≤ 2.0mmol/l





Medications for Hyperlipidemia

Statins

Generic Name	Trade Name
Atorvastatin	Lipitor
Fluvastatin	Lescol
Lovastatin	Mevacor, generic
Pravastatin	Pravachol, generic
Rosuvastatin	Crestor, generic
Simvastatin	Zocor, generic





Medications for Hyperlipidemia

Other Medications

- PCSK-9
- Bile acid seqestrants
- Cholesterol absorption inhibitors
- Fibrates
- Nicotinic acid





Medication Changes during Illness

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



Sick Day Medication Guideline



Prevent hypoglycemia

- Adjust insulin
 - Test more often
- Stop secretagogues

Prevent dehydration



Sick Pay Medication Guideline

Medications that affect kidney function when dehydrated

- ACE
- ARB
- Diuretics
- Metformin
- SGLT2 inhibitors
- Anti-inflammatory pain medication

If the symptoms last more than 24 hours and you continue to be dehydrated, or at risk of dehydration, you should also TEMPORARILY STOP:

Certain Blood Pressure / Heart Medications

- ACE Inhibitors: e.g. Enalapril (Vasotec*), Fosinopril (Monopril*), Lisinopril (Prinivil*/Zestril*), Perindopril (Coversyl*), Quinapril (Accupril*), Ramipril (Altace*), Trandolapril (Mavik*)
- ARBs: e.g. Candesartan (Atacand®), Eprosartan (Teveten®), Irbesartan (Avapro®), Losartan (Cozaar®), Olmesartan (Olmetec®), Telmisartan (Micardis®), Valsartan (Diovan®)

All Water Pills

 e.g. Chlorthalidone (Hygroton), Furosemide (Lasix[®]), Hydrochlorothiazide, Indapamide (Lozide[®]), Metolazone (Zaroxolyn[®]), Spironolactone (Aldactone[®])

Certain Diabetes Pills

- . Metformin (Glucophage® or Glumetza®)
- SGLT2 Inhibitors: e.g. Canagliflozin (Invokana®), Dapagliflozin (Forxiga®), Empagliflozin (Jardiance™)

Anti-Inflammatory Pain Medications

 e.g. Ibuprofen (Advil®/Motrin®), Celecoxib (Celebrex®), Diclofenac (Voltaren®), Ketorolac (Toradol®), Napoxen (Aleve®/Naprosyn®)

Note: The list above does not include the names of medications that come in combination (2 medications in one tablet).

Ask your pharmacist to tell you:

The medications I need to TEMPORARILY STOP are:

When I am eating less than normal:

When I am dehydrated:

This personalized list last reviewed (date):

Note: RESTART these medications when you are eating and drinking normally.

Call your health-care team (Pharmacist, Doctor, Nurse Practitioner, Nurse, Dietitian) and/or go the Emergency Department

- . If you cannot drink enough fluids
- . If you don't know which medications to stop
- . If you don't know how to adjust your insulin
- If you have been told to check your ketones and they are moderate to high
- If you have any of the following that are not getting better: vomiting, diarrhea, stomach pain, frequent urination, extreme thirst, weakness, difficulty breathing or fever



- Seema is presently on glimipride and metformin. Acarbose has been added as the A1C is still elevated.
- What would be the most important information to tell her about this change in medication?
- a) Acarbose does not cause hypoglycemia
- b) Fruit juice is the best way to treat hypoglycemia
- Hypoglycemia must be treated with glucose tablets or milk
 - d) Hypoglycemia is best treated with food e.g. crackers
 - e) If hypoglycemia occurs the metformin should be reduced.



- Alfred is 75 and has recently returned home after an MI. His eGFR is 60. His present medications include glucophage 1 gm bid, lisinopril 40 mg od, atorvastatin 20 mg. He develops flu-like symptoms and is vomiting. What would you tell him about his medications?
- a) Stop all medications as illness will decrease blood glucose
- b) Stop lisinopril and atorvastatin
- Stop glucophage and lisinopril
 - d) Continue with all medications



Paula calls her diabetes educator as she has had 2 genital mycotic infections in the last two months. She reports that her blood sugars are in good control with FBS 5.6-7.1mmol/l and postprandial sugars all under 10 mmol/l. Her medications include glucophage 1 gm bid, repaglinide 1 mg tid and canagliflozin 100 mg, candesartan 16 mg. What is the most likely explanation?

- a) Glucophage can cause dehydration
- There is an increased risk of genital infections with canagliflozin
 - c) Repaglinide and glucophage should not be taken together
 - d) Repaglinide can cause dehydration and risk of genital infections



Nasar (age 39) has had type 2 diabetes for 2 years. He has recently immigrated to Canada. His A1c is 8.4% and eGFR 110. He is on glucophage 1g bid. What class of medication would you recommend adding given his limited finances and no drug coverage.

- a) DPP4
- b) SGLT2
- c) GLP-1





Nikki questions whether her blood glucose meter is working correctly. You send her for a lab to meter comparison. According to the 2018 guidelines what should the correlation be?

- a) 5 %
- b) 10%
- 15%
 - d) 20%



Nikki comes to the clinic reporting her period is overdue by 4 weeks. Which medications should be stopped?

- a) Vitamin D + omega 3
- Lovastatin + olmestartan
 - c) Metformin
 - d) Folic acid



What statement is true about alternate site testing?

- a) It can be done at any time
- b) The forearm is the best place to test right after a meal
- The base of the thumb is most comparable with fingertip testing
- d) It should only be used by children



What would be a realistic expectation for reducing the progression to diabetes with Metformin?

- a) 20 %
- 31%
 - c) 58%
 - d) 10%



Questions



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