



Taddle Creek

Family Health Team

Title:	Diabetes Management in Adults	Number:	TCFHT-MD05
Activation Date:	12-June-2012	Review Date:	25-July-2022
Next Review Date:	12-June-2023		

Sponsoring/Contact Person(s)
(name, position, contact particulars):

Karen Finch, Registered Nurse, Certified Diabetes Educator
790 Bay St. Suite 508
Toronto, Ontario M5G 1N8
416- 204-1256, x502

Dr. Mitch Vainberg
790 Bay Street, Suite 300
Toronto, Ontario M5G 1N8
416-960-1366

Sherry Kennedy, Executive Director – skennedy@tcfht.on.ca
790 Bay Street, Suite 306,
Toronto, Ontario M5G 1N8
416-260-1315, x307

Order and/or Delegated Procedure: Appendix Attached: No Yes

Title: Appendix C – Performed Controlled Acts and Procedures

The implementers are authorized to perform the following activities, in accordance with Appendix C when all conditions in this directive and the attached appendices are met:

- 1) Start, Adjust, Hold and Discontinue Basal Insulin already prescribed by Primary Care Provider/Endocrinologist
- 2) Start, Adjust, Hold and Discontinue Intensive Insulin Therapy already prescribed by Primary Care Provider/Endocrinologist
- 3) Adjust, Hold and Discontinue Oral Antihyperglycemic Agents
- 4) Adjust, Hold and Discontinue Injectable Antihyperglycemic Agents
- 5) Dispense Insulin prescribed by Primary Care Provider or Endocrinologist

Recipient Patients: Appendix Attached: No Yes

Title: Appendix A – Authorizer Approval Form

Recipients must:

- Be active patients of a TCFHT primary care provider/endocrinologist who has approved this directive by signing the Authorizer Approval Form
- Have a diagnosis of diabetes mellitus (type 1 or type 2) or pre-diabetes
- Meet the conditions identified in this directive

Authorized Implementers:**Appendix Attached:** No Yes**Title:** Appendix B – Implementer Approval Form; Appendix D – Competency Checklist

Implementers must be TCFHT employed Regulated Health Care Providers or Physician Assistant (under the supervision of a physician).

Implementers must complete the following preparation and sign the Implementer Approval Form:

- Must become a certified diabetes educator (CDE) as per the Canadian Diabetes Educator Certification Board
- Practice according to Diabetes Canada's (DC's) most current Clinical Practice Guidelines
- Assess their own knowledge, skill, and judgment to competently perform these directives

In addition, to implement delegated procedures 1-4 (start, adjust, hold or discontinue basal or intensive insulin therapy to adjust, hold and discontinue oral antihyperglycemic agents, and to adjust, hold and discontinue injectable antihyperglycemic agents):

- Must have at least 1600 practice hours in providing direct diabetes education with patients living with diabetes
- Must be mentored by an authorized implementer, demonstrate the competencies and review all of the DC's Best Practice Guidelines, as outlined in Appendix D
- Must be supervised by Endocrinologist with at least 3 patient cases involving adjustments to insulin, oral antihyperglycemic agents, and injectable antihyperglycemic agents and complete and sign competency performance checklist (Appendix D)

Indications: **Appendix Attached:** No Yes**Title:** Appendix C – Performed Controlled Acts and Procedures

- In general, each action/procedure under each directive will be implemented in the context of the existing physician/nurse practitioner relationship and as part of the medical diagnosis and plan of care established by the physician/nurse practitioner. These actions/procedures will be implemented without specific prior discussion (but as part of the plan of care) as per the indications and contraindications for each of these directives (Appendix C).
- Communication to primary care provider/endocrinologist regarding medication/insulin adjustments made following implementation of medical directive (see documentation and communication)

Contraindications:

- Indications described in Appendix C are not met

<p>Consent: Appendix Attached: <input checked="" type="checkbox"/> No <input type="checkbox"/> Yes</p> <p>Title:</p> <ul style="list-style-type: none"> • Patient’s consent is implied, as patient has presented seeking support with diabetes management, and is a Family Health Team patient, where interprofessional practice is expected • The implementer fully explains potential risks and benefits prior to initiating/adjusting insulin therapy and with any changes to oral or injectable antihyperglycemic medications
<p>Guidelines for Implementing the Order/Procedure: Appendix Attached: <input type="checkbox"/> No <input checked="" type="checkbox"/> Yes</p> <p>Title: Appendix C – Performed Controlled Acts and Procedures As per Appendix C.</p>
<p>Documentation and Communication: Appendix Attached: <input checked="" type="checkbox"/> No <input type="checkbox"/> Yes</p> <p>Title:</p> <ul style="list-style-type: none"> • Documentation in the patient’s eMR needs to include: name and number of the directive, patient’s Capillary Blood Glucose (CBG) and/or Flash Blood Glucose (FBG) patterns (if applicable), current medications, self-management skills and learning needs, clinical findings and the plan of care, patient’s response to the procedure or directions provided • Implementer will send a message in Practice Solutions to patient’s primary care provider, notifying him/her that patient was seen, recommendations were made and that eMR note needs review for details. • Primary care provider is responsible for entering newly prescribed medications into medication list in Practice Solutions. Implementer is responsible for updating medication list in eMR and informing the patient’s pharmacy via fax and/or phone call. • Implementer will arrange a follow up plan in collaboration with patient and primary care provider/endocrinologist with any medication/insulin adjustments.

Review and Quality Monitoring Guidelines:Appendix Attached: No Yes

Title: Appendix D – Competency Checklist

- Routine review will occur annually on the anniversary of the activation date. Review will involve a collaboration between the authorizing primary care providers and the authorized implementers.
- At any such time that issues related to the use of this directive are identified, TCFHT must act upon the concerns and immediately undertake a review of the directive by the authorizing primary care providers and the authorized implementers.
- If new information becomes available between routine renewals, such as the publishing of new Diabetes Canada Best Practice Guidelines, and particularly if this new information has implications for unexpected outcomes, the directive will be reviewed by the authorizing primary care providers and the authorized implementers.
- This medical directive can be placed on hold if routine review processes are not completed, or if indicated for an ad hoc review. During the hold, implementers cannot perform the procedures under authority of the directive and must obtain direct, patient-specific orders for the procedure until it is renewed.
- **Implementer’s competencies will be reviewed on a yearly basis as part of their performance review. Clinicians must consistently demonstrate competency in order to remain an authorized implementer of this directive – see Appendix D**

References:

Diabetes Canada. (2010). Building Competency in Diabetes Education: Advancing Practice.

Diabetes Canada. (2020). Building Competency in Diabetes Education: The Essentials (5th Edition).

Diabetes Canada. (2018). Clinical Practice Guidelines for the Prevention and Management of Diabetes in Canada. Retrieved from <http://guidelines.diabetes.ca/cpg>

Diabetes Canada. (2020). Pharmacologic Glycemic Management of Type 2 Diabetes in Adults: 2020 Update. Retrieved from <http://guidelines.diabetes.ca/cpg/chapter-13-2020-update>

Canadian Insulin Injection Recommendations: FIT Technique Plus, 2020. Retrieved from <http://www.fit4diabetes.com/canada-english/fit-technique-plus/>

Fit Forum for Injection Technique Canada. (2020). Fit Forum for Injection Technique Canada: Recommendations for Best Practice in Injection Technique 4th Edition. Retrieved from <http://fit4diabetes.com/canada-english/fit-recommendations/>

Health Canada. Drug Product database. Retrieved from <https://www.canada.ca/en/health-canada/services/drugs-health-products/drug-products/drug-product-database.html>

Ontario College of Family Physicians Insulin Prescription Tool – December 2017. Retrieved from <https://www.diabetes.ca/DiabetesCanadaWebsite/media/Managing-My-Diabetes/Tools%20and%20Resources/insulin-prescription-fillable-EN.pdf?ext=.pdf>

UpToDate. Retrieved from <https://www.uptodate.com>

NOTE:

This medical directive is based on TCFHT’s previous medical directive DEP1 entitled, “Diabetes Management in Adults Medical Directive,” which required revision in formatting to reflect the growth of the TCFHT organization. The majority of the content of DEP1 has remained the same for the revised TCFHT-MD05 version. Therefore, all approved Implementers and Authorizers for medical directive DEP1 “Diabetes Management in Adults Medical Directive” have grandfathered approval for TCFHT-MD05 “Diabetes Management in Adults.”

Appendix A:

Authorizer Approval Form

Name	Signature	Date
<hr/>	<hr/>	<hr/>
<hr/>	<hr/>	<hr/>
<hr/>	<hr/>	<hr/>
<hr/>	<hr/>	<hr/>
<hr/>	<hr/>	<hr/>
<hr/>	<hr/>	<hr/>
<hr/>	<hr/>	<hr/>
<hr/>	<hr/>	<hr/>
<hr/>	<hr/>	<hr/>
<hr/>	<hr/>	<hr/>
<hr/>	<hr/>	<hr/>
<hr/>	<hr/>	<hr/>
<hr/>	<hr/>	<hr/>
<hr/>	<hr/>	<hr/>
<hr/>	<hr/>	<hr/>
<hr/>	<hr/>	<hr/>
<hr/>	<hr/>	<hr/>
<hr/>	<hr/>	<hr/>
<hr/>	<hr/>	<hr/>
<hr/>	<hr/>	<hr/>
<hr/>	<hr/>	<hr/>

Appendix B:

Implementer Approval Form

To be signed when the implementer has completed the required preparation, and feel they have the knowledge, skill, and judgement to competently carry out the actions outlined in this directive.

Delegated Procedures 1-4:

- Start, Adjust, Hold and Discontinue Bedtime Insulin
- Start, Adjust, Hold and Discontinue Intensive Insulin Therapy
- Adjust, Hold and Discontinue Oral Antihyperglycemic Agents
- Adjust, Hold and Discontinue Injectable Antihyperglycemic Agents

Name	Signature	Date
_____	_____	_____
_____	_____	_____
_____	_____	_____
_____	_____	_____
_____	_____	_____
_____	_____	_____

Delegated Procedures 5:

- Dispense Insulin or Injectable antihyperglycemics prescribed by Primary Care Provider or Consulting Endocrinologist

Name	Signature	Date
_____	_____	_____
_____	_____	_____
_____	_____	_____
_____	_____	_____
_____	_____	_____

Appendix C

Performed Controlled Acts and Procedures

Table 1: List of Medications (Insulin) Implemented under this Directive with Detailed Indications/Contraindications

Insulin Type	Onset	Peak/ Duration	Indications for Adjustment	Therapeutic Considerations
Rapid-Acting <ul style="list-style-type: none"> • Lispro U-100, U-200 (Humalog) • Aspart (NovoRapid) • Glulisine (Apidra) • Aspart (Fiasp) 	2-15 minutes	1-2 hours/ 3-5 hours	<ul style="list-style-type: none"> • 2 hr post-meal and/or pre-meal blood sugars (lunch, dinner) are either elevated or hypoglycemia occurs 	<ul style="list-style-type: none"> • For current basal insulin users, maintain the basal dose and add bolus insulin with each meal at a dose equivalent to 10% of the basal dose. For example, if the patient is on 50 units of basal insulin, add 5 units of bolus insulin with each meal. • For new insulin users starting a full Basal + Bolus regimen, calculate Total Daily Insulin dose (TDI) as 0.3 to 0.5 units/kg, then distribute as follows: – 40% of TDI dose as basal insulin (Lantus, Levemir, Toujeo™, Humulin N, Novolin ge NPH) at bedtime. – 20% of TDI dose as prandial (bolus) insulin prior to each meal. • Patients should eat within 0-15 minutes after injection • In certain circumstances injecting shortly after eating is appropriate e.g. GI disturbance resulting in early satiety, vomiting etc. • For Patients on MDI (multiple daily injections) using a rapid-acting insulin it is recommended that they self-monitor blood glucose (SMBG) at least QID for safe titration of doses • Humalog 200 must only be used in the pre-filled pen. It must not be drawn up in a syringe • Fiasp is best injected 2 minutes before and up to 20 minutes after a meal. It may not be best in people with delayed gastric emptying & has not been studied in pregnancy

Insulin Type	Onset	Peak/ Duration	Indications for Adjustment	Therapeutic Considerations
Short-Acting <ul style="list-style-type: none"> • Humulin R • Novolin Toronto 	30 minutes	2-3 hrs/6.5 hours	<ul style="list-style-type: none"> • Pre-meal blood sugars (lunch, supper) and/or bedtime are elevated or hypoglycemia occurs 	<ul style="list-style-type: none"> • Same initiation recommendations as rapid-acting insulin (see above) • Rapid-acting insulin usually preferable to short-acting due to better post-prandial glycemic control and lower risk for hypoglycemia • Patients should inject insulin 30 minutes prior to eating • For Patients on MDI (multiple daily injections) using a short-acting insulin it is recommended that they SMBG at least QID for safe titration of doses
Intermediate-Acting <ul style="list-style-type: none"> • Humulin N • Novolin NPH 	1-3 hours	5-8 hours/up to 18hours	<ul style="list-style-type: none"> • Increase evening dose if high fasting blood sugar (if rebound hyperglycemia has been eliminated) • Decrease dose if hypoglycemia (recommend CBG testing in the night to rule out nocturnal hypoglycemia) • Increase or decrease morning dose if 5-8 hour pre-meal blood sugars are out of target 	<ul style="list-style-type: none"> • To be injected once to twice daily in the morning and/or before bed • Start at 10u daily or 0.2u/kg • Long-acting basal insulins are typically preferred due to longer action profile and lower risk for hypoglycemia but intermediate-acting may be appropriate in some cases e.g., affordability/, glycemic management of hyperglycemia due to steroid therapy etc. • Must be adequately re-suspended before injecting • Recommended for Patients to SMBG OD-BID for safe titration of doses

Table 1 Notes:

- The implementer will adhere to the indications and contraindications outlined in Table 1
- The implementer is responsible for teaching patient safe injection technique according to FIT 2020 guidelines.
- A prescription is required from the primary health care provider and/or endocrinologist prior to insulin initiation.
- Individual considerations need to be assessed with insulin initiation i.e. patients who are hypoglycemia unaware or have a fear of insulin-induced hypoglycemia can be initiated on a smaller dose etc.
- Evidence-based recommendations are to adjust insulin by 1-2u q 3-4 days or by 1u per day. Under certain circumstances patients may need insulin adjustment greater or less than 5-10% of total daily dose i.e. extreme hyperglycemia, medications and/or lifestyle factors that can increase/decrease glycemic levels.
- Implementer should determine a communication plan with the patient for further insulin adjustment and encourage self-titration if possible.
- Implementer will instruct a patient starting insulin around hypoglycemia treatment, driving instructions and instruct family/caregivers on using a glucagon kit when applicable i.e. type 1 diabetes and high risk for hypoglycemia.
- In the case of high CBG readings and low CBG readings, always correct for hypoglycemia first.
- Patients should be instructed as to how to adjust insulin during times of illness, travel and physical activity following current best practice guidelines.
- Allergic reactions are rare but can occur with a few patients. Reactions may be local (i.e. rash/weal at site) or systemic (i.e. shortness of breath, wheezing or severe weakness). Implementer should instruct Patient to hold insulin and get in contact with primary care provider a.s.a.p. and/or proceed to the nearest emergency department.
- Patient should be instructed to store unopened insulin vials/cartridges in the refrigerator, store open insulin vials/cartridges at room temperature, not expose insulin to heat or direct sunlight, not freeze insulin, and to use by expiration date.
- The primary care provider or endocrinologist must be available to provide consultation as required.
- Primary care provider and/or endocrinologist should be consulted in the following circumstances:
 - Recurrent or severe hypoglycemia with no apparent cause
 - Glycemic control is not improving or is deteriorating despite adjustments made to insulin or other component of the treatment plan
 - Total daily dose exceeds what is generally expected for age/body type
 - Patient shows signs and symptoms of Diabetic Ketoacidosis (DKA), dehydration or other serious problems *send to the Emergency Department immediately
 - Recurring/persistent vomiting/diarrhea
 - Disordered eating pattern resulting in calorie restriction
 - Significant error in dose or timing of insulin administered by person or caregiver

- Change in brand or type of insulin
- Change in frequency of injections i.e., BID to TID
- For patients with additional complex medical or endocrine disorders which may influence insulin requirements or patient safety
- In all situations that are beyond the implementer's scope of practice and/or competency level

Table 2: Non-Insulin Antihyperglycemic agents: List of Medications Implemented under this Directive with Detailed Indications/Contraindications

Antihyperglycemic Agent	Indications for Adjustment	Contraindications/Precautions
<p>Alpha-Glucosidase Inhibitor</p> <p>Acarbose (Glucobay, Prandase)</p> <p><i>Initial Dose:</i> 25-50 mg daily</p> <p><i>Average Dose (Max dose):</i> 50-100 mg tid (300 mg)</p> <p><i>Special Instructions:</i> Take with first bite of meal</p> <p><i>Onset/peak/duration:</i> 1h/2h/4-6h</p> <p><i>Expected HbA1C reduction:</i> 0.7-0.8%</p> <p>ODB Coverage: limited use</p>	<ul style="list-style-type: none"> • Gastrointestinal (GI) side effects • Inadequate blood glucose control • Very low frequency of hypoglycemia unless combined with a sulfonylurea • Initiate therapy with 25mg OD-BID and titrate slowly by 25mg/day every 2-4 weeks as tolerated • Maximum effectiveness with at 50mg TID; higher doses associated with increased adverse events 	<ul style="list-style-type: none"> • Not recommended as initial therapy in people with severe hyperglycemia (A1C \geq8.5%) • Gastrointestinal side effects in approx. 30% of Patient's i.e. cramps, diarrhea, abdominal distension, flatulence (effects usually decrease with continued use but there is a high discontinuation rate based on GI side effects) • Treat hypoglycemia with Dextrose tablets, milk or honey as Acarbose interferes with glucose absorption • Contraindicated in Patient's with DKA, inflammatory bowel disease, intestinal ulcers, cirrhosis, partial intestinal blockage or predisposed to blockage • Renal dosing: discontinue use if creatinine clearance (CrCl) <25ml/min or eGFR <30 mL/min • In patients with known liver impairment or liver disease, liver enzymes should be monitored prior to start of Acarbose, and monitored on a regular basis within the first year • Case reports of reduction in absorption of digoxin and increased absorption of warfarin Maximum doses based on weight <132 lbs: 50mg TID; >132 lbs: 100mg TID

Antihyperglycemic Agent	Indications for Adjustment	Contraindications/Precautions
<p>Biguanides</p> <p><u>Metformin (Glucophage)</u> <i>Initial Dose:</i> 250-500 mg daily <i>Average Dose (Max dose):</i> 500-1000 mg bid or 850 mg tid (2550 mg) <i>Special Instructions:</i> Take with meals to reduce GI side effects <i>Onset/peak/duration:</i> 1-2h/6h/6-12h <i>Expected HbA1C reduction:</i> 1.0% ODB Coverage: Yes</p> <p><u>Glumetza (Metformin HCL ER)</u> <i>Initial Dose:</i> 500 mg daily (ideally with dinner) <i>Average Dose (Max dose):</i> 1000-2000 mg daily (2500 mg) <i>Onset/peak/duration:</i> 1-2h/4-8h/17.6-19.8h <i>Expected HbA1C reduction:</i> 1.0% ODB Coverage: No</p>	<ul style="list-style-type: none"> • Initiate Glucophage 500mg twice daily or 850mg OD • Initiate Glumetza 500mg once or twice daily • GI side effects in 20-30% of Patient's (Glumetza associated with fewer GI side effects than short-acting Metformin). Side effects can be reduced by slow titration (500mg/day every 2 weeks and taking medication with meals) • Decrease in FBG levels seen within 3-5 days; maximal effect in 1-2 weeks • 80-85% of glucose lowering effect is seen with 1500mg/day • Maximum effective dose is 2000mg/day • Renal insufficiency • Hypoglycemia (rare as monotherapy) • Blood glucose remains above target • Biguanides should be put on hold in patients with severe dehydration (i.e., vomiting and unable to keep down fluids) 	<ul style="list-style-type: none"> • Contraindicated in people with a history of lactic acidosis, severe hepatic dysfunction, severe infection/dehydration, trauma or cardiorespiratory insufficiency, surgery or in excessive alcohol intake, whether acute or chronic • Reduced dose recommended if CrCl/eGFR <45mL/min and contraindicated if CrCl/eGFR <30 mL/min due to risk of accumulation • Glucophage should not be initiated if eGFR 30-44, but can be maintained • 5-10% of people are unable to tolerate due to substantial GI side effects (upset stomach, nausea, diarrhea, anorexia, metallic taste) • Metformin should be stopped during acute illness (severe infections, trauma, surgery) and the recovery phase afterwards. Should also be put on hold in patients with severe dehydration (i.e., vomiting and unable to keep down fluids) • Should do baseline liver function tests (LFT's) • Higher doses (above 2000mg/day) associated with increased risk of adverse events with no additive effect • Hold for 48 hours if undergoing radiologic studies with administration of iodinated contrast material (hold on day of procedure until 2-3 days after) • Not recommended in the elderly (over 80yrs) unless CrCl/eGFR is >60 mL/min due to decreased muscle mass • Recommend conservative dosing in the elderly • Lactic acidosis is rare 0.03/1000 patients and 0.015 fatal cases/1000 patients; more likely to occur in patients with renal insufficiency, alcohol or liver disease. Hold dose in hypoxic states, shock, severe infection or septicemia • Measurements of serum vitamin B12 are advisable at least every 1 to 2 years in patients on long-term treatment (Product Monograph – Health Canada)

<p>Insulin Secretagogues Sulfonylureas:</p> <p><u>Diamicron (Gliclazide)</u> <i>Initial Dose:</i> 40-80 mg daily or bid with meals <i>Average Dose (Max dose):</i> 80-160 mg bid (320 mg) <i>Onset/peak/duration:</i> 1-2h/4-6h/10-14h <i>Expected HbA1C reduction:</i> 0.6-1.2% ODB Coverage: Yes</p> <p><u>Diamicron MR (Gliclazide MR)</u> <i>Initial Dose:</i> 30 mg daily with first meal <i>Average Dose (Max dose):</i> 30-120 mg daily (120 mg) <i>Onset/peak/duration:</i> 1-2h/6-12/>24h <i>Expected HbA1C reduction:</i> 0.6-1.2% ODB Coverage: Yes</p> <p><u>Diabeta (Glyburide)</u> <i>Initial Dose:</i> 2.5 mg – 5.0 mg daily or bid with meals <i>Average Dose (Max dose):</i> 5-10 mg bid with meals (20 mg) <i>Special Instructions:</i> Take 30min prior to meal) <i>Onset/peak/duration:</i> 1-2h/4-6h/10-14h <i>Expected HbA1C reduction:</i> 0.6-1.2% ODB Coverage: Yes</p> <p><u>Amaryl (Glimepiride)</u> <i>Initial Dose:</i> 1 mg daily with first meal <i>Average Dose (Max dose):</i> 1-4 mg daily (8 mg) <i>Onset/peak/duration:</i> 20min/2-4h/24h <i>Expected HbA1C reduction:</i> 0.6-1.2% ODB Coverage: No</p>	<ul style="list-style-type: none"> • Frequent hypoglycemia (decrease or discontinue if hypoglycemia persists 1-2 times per week) • Inadequate blood glucose control (blood glucose remains above target consistently) • Dose should be started low and titrated every 1-2 weeks until glycemic targets are met • Sulfonylureas should be put on hold in patients with severe dehydration (i.e., vomiting and unable to keep down fluids) 	<ul style="list-style-type: none"> • Associated with weight gain (unless dietary modifications are made) • Associated with hypoglycemia; annual rate of any hypoglycemia is 20%. Major hypoglycemic events occur in 1-2 % of individuals • Consider using other class(es) of oral antihyperglycemic agents first in patients at high risk of hypoglycemia i.e. the elderly • Requires lower dose and slower titration in patients with hepatic/renal impairment and the elderly • Increased risk for hypoglycemia with insulin • Glyburide not recommended with eGFR <60 mL/min. • Gliclazide and Glimepiride should be used with caution if eGFR 30-59 mL/min, not recommended with eGFR <30 mL/min
------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------	------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------	---------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------

Antihyperglycemic Agent	Indications for Adjustment	Contraindications/Precautions
<p>Insulin Secretagogues Non-Sulfonylureas: Meglatinides</p> <p><u>Gluconorm (Repaglinide)</u> <i>Initial Dose:</i> 0.5-1mg tid with meals <i>Average Dose (Max dose):</i> 0.5-4 mg tid (16 mg) <i>Special Instructions:</i> take 1-30min before meals <i>Onset/peak/duration:</i> 30min/1h/4-5h <i>Expected HbA1C reduction:</i> 0.7-1.1% ODB Coverage: Exceptional Access Program (EAP)</p>	<ul style="list-style-type: none"> • Frequent hypoglycemia (decrease or discontinue if hypoglycemia persists 1-2 times per week) • Less hypoglycemia compared to sulfonylureas and are ideal for patients with irregular meal times • Inadequate blood glucose control (blood glucose remains above target consistently) • Doses should be titrated weekly as required to obtain glycemic targets 	<ul style="list-style-type: none"> • Less likely to cause weight gain and hypoglycemia than sulfonylureas • Safe to use in renal impairment and mild hepatic impairment but requires slower dose titration • In the elderly Repaglinide should be initiated at 0.5mg TID and titrate dose slowly (especially with CrCl 20-39mL/min) • Preferred for use in elderly individuals with erratic eating patterns • The concomitant use of Repaglinide and Clopidogrel (Plavix) is contraindicated as it may lead to a significant decrease in blood glucose levels due to a drug-drug interaction

<p>Thiazolidinediones (TZD's)</p> <p><u>Actos (Pioglitazone)</u> <i>Initial Dose:</i> 15 mg daily <i>Average Dose (Max dose):</i> 15-45 mg daily (45 mg) <i>Onset/peak/duration:</i> 30min/2-4h/4 weeks <i>Expected HbA1C reduction:</i> 0.7-0.9% ODB Coverage: EAP</p> <p><u>Avandia (Rosiglitazone)</u> <i>Initial Dose:</i> 4 mg daily <i>Average Dose (Max dose):</i> 2-8 mg daily (8 mg) <i>Onset/peak/duration:</i> 30-60min/1-2h/4 weeks <i>Expected HbA1C reduction:</i> 0.7-0.9% ODB Coverage: EAP</p>	<ul style="list-style-type: none"> • Edema • Shortness of breath • Discontinue TZD if insulin is initiated • Titrate every 2-4 weeks • Full BG-lowering effect seen within 6-12 weeks • Discontinue if ALT >3 X upper limit of normal 	<ul style="list-style-type: none"> • Should do baseline LFT's prior to initiation, every 2 months for the first year and then periodically • May induce edema, fluid retention (recommended to monitor weight) • Pioglitazone may increase risk of bladder cancer and is not recommended for use with Patient's who have or have had bladder cancer, are at high risk, have blood or a red color in their urine. Patient's taking pioglitazone should be assessed regularly for potential symptoms of bladder cancer i.e. blood or red color in urine, painful urinate etc. • Can cause weight gain (subcutaneous fat + fluid retention) 1.5-4.84kg but decrease in visceral and hepatic fat. Weight gain is generally dose dependent • Associated with increased risk for bone loss and fractures in women 55yrs or older (0.78 per 100) • Used in combination with insulin may increase risk of edema and CHF. The combination of a TZD plus insulin is currently not an approved treatment in Canada • Rosiglitazone is no longer approved for use alone to treat DM2 except when Metformin use is contraindicated or not tolerated and all other oral agents have been tried alone or together and targets are not reached • Rosiglitazone is not indicated for triple therapy and is only indicated in combination with Metformin or a sulfonylurea • Pioglitazone is no longer approved for use with Metformin and a sulfonylurea • Can be used safely in mild to severe renal impairment • Contraindicated for people with CHF New York Heart Association (NYHA) Class I to IV cardiac status; evidence of left ventricular dysfunction or serious hepatic impairment (ALT 2.5 X upper limit of normal) • Prior to prescribing Avandia practitioners must 1) document the individual's eligibility to meet the above criteria; 2) counsel the individuals on the risks and benefits of Avandia, including the CV risks; and 3) obtain the person's written informed consent to take the drug
--------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------	------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------	-----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------

<p>DPP4 Inhibitors</p> <p><u>Sitagliptin (Januvia)</u> <i>Initial Dose:</i> 100 mg daily qam with/without food <i>Average Dose (Max dose):</i> 100 mg daily (100 mg) <i>Onset/peak/duration:</i> Rapidly absorbed/1-4h/24h <i>Expected HbA1C reduction:</i> 0.5-0.7% ODB Coverage: Yes</p> <p><u>Saxagliptin (Onglyza)</u> <i>Initial Dose:</i> 5 mg daily with/without food <i>Average Dose (Max dose):</i> 5 mg daily (5 mg) <i>Onset/peak/duration:</i> Rapidly absorbed/2.5h/26.9h <i>Expected HbA1C reduction:</i> 0.5-0.7% ODB Coverage: Yes</p> <p><u>Linagliptin (Trajenta)</u> <i>Initial Dose:</i> 5 mg daily with/without food <i>Average Dose (Max dose):</i> 5 mg daily (5 mg) <i>Onset/peak/duration:</i> Rapidly absorbed/1.5h/24h <i>Expected HbA1C reduction:</i> 0.5-0.7% ODB Coverage: Yes</p> <p><u>Alogliptin (Nesina)</u> <i>Initial Dose:</i> 25 mg daily with/without food <i>Average Dose (Max dose):</i> 25 mg daily (5 mg) <i>Onset/peak/duration:</i> Rapidly absorbed/1-2h/approx. 24h <i>Expected HbA1C reduction:</i> 0.5-0.7% ODB Coverage: No</p>	<ul style="list-style-type: none"> • Nasopharyngitis, cough and headache (rare cases) • Severe joint pain (rare cases), usually within 1 month of initiation • Inadequate glucose control • Increased risk for hypoglycemia if combined with a sulfonylurea • Discontinue if suspicion of pancreatitis i.e. severe ongoing stomach or back pain with/without vomiting 	<ul style="list-style-type: none"> • Linagliptin can be used in renal insufficiency (eGFR <15 mL/min and dialysis) • Saxagliptin dose should be decreased to 2.5 mg od if eGFR <50 mL/min, and discontinued if eGFR <15 mL/min. It should not be used in patients on dialysis (assess renal function prior to treatment and periodically after) • Sitagliptin dose should be decreased to 50 mg od if eGFR is 30-49 mL/min, and decreased further to 25 mg od if eGFR <30 mL/min (assess renal function prior to treatment and periodically after) • Alogliptin dose should be decreased to 12.5 mg if eGFR is 30-59 mL/minute, 6.25 mg od if eGFR <30 mL/ and can be used at 6.25mg in ESRD with hemodialysis (has not been studied with peritoneal dialysis) • Use in caution in the elderly (as per renal guidelines) • Safety profile has not been studied and is unclear in individuals who are immunocompromised e.g. lymphocyte abnormalities, HIV, or people who have undergone organ transplant • Approved for use with Metformin and a sulfonylurea • Approved for use with insulin except for Linagliptin • Linagliptin, Sitagliptin and Alogliptin are not recommended in severe hepatic insufficiency and Saxagliptin is not recommended in moderate to severe hepatic impairment (monitor hepatic function before initiating treatment and periodically after) • Not recommended if history of pancreatitis • Caution if history of alcoholism, high triglycerides (higher risk for pancreatitis) • Not recommended for people with heart failure • Contraindicated for use for patients with personal or family history of medullary thyroid carcinoma or in patients with Multiple Endocrine Neoplasia syndrome 2
---------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------	--------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------	-----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------

Antihyperglycemic Agent	Indications for Adjustment	Contraindications/Precautions
<p>GLP-1 Receptor Agonist (oral)</p> <p><u>Semaglutide (Rybelsus)</u></p> <p><i>Initial Dose:</i> 3 mg daily without food</p> <p><i>Average Dose (Max dose):</i> 7 mg daily (14 mg)</p> <p><i>Onset/peak/duration:</i> rapidly absorbed/1 hour/approx 1 week</p> <p><i>Expected HbA1C reduction:</i> 0.6-1.4%</p> <p>ODB Coverage: No</p>	<ul style="list-style-type: none"> • Gastrointestinal (GI) side effects • Inadequate blood glucose control • low frequency of hypoglycemia unless combined with a sulfonylurea or insulin • Initiate therapy with 3mg OD. After 30 days, increase to a maintenance dose of 7mg once daily. If additional glycemic control is needed after at least 30 days on the 7 mg dose, the dose can be increased to a maintenance dose of 14mg once daily. This regimen is intended to mitigate gastrointestinal symptoms during dose escalation • Regular self-monitoring of blood glucose is not needed in order to adjust the dose of Rybelsus on its own. However, when used in combination with a Sulfonylurea or insulin, SMBG testing may be necessary in order to reduce the risk of hypoglycemia 	<ul style="list-style-type: none"> • Most frequent adverse reactions include: nausea, abdominal pain, diarrhea and vomiting • Rybelsus must be taken on an empty stomach at least 30 min before the first food, beverage or other oral medications of the day. Waiting less than 30 min is likely to decrease the amount of semaglutide absorbed • Rybelsus should be taken with more than half a glass of water equivalent to 120 mL. A larger volume of water is likely to decrease the amount of Semaglutide absorbed. • Rybelsus should be swallowed whole. Do not split, crush or chew • If a dose is missed, the missed dose should be skipped, and the next dose should be taken the following day • Contraindicated for use for patients with personal or family history of medullary thyroid carcinoma or in patients with Multiple Endocrine Neoplasia syndrome 2 • Rybelsus should not be used for patients with Type 1 Diabetes or in DKA or in pregnancy • Women of childbearing potential are recommended to use contraception when treated with Semaglutide. If a patient wishes to become pregnant or pregnancy occurs. Rybelsus should be discontinued. Rybelsus should be discontinued at least 2 months before a planned pregnancy due to the long half-life of Semaglutide • Caution for use with those with cardiac conditions that might be worsened by an increased in HR (i.e. tachyarrhythmias) • Semaglutide causes a prolongation of the PR interval, and caution should be observed in patients with pre-existing conduction system abnormalities (first/second/third degree AV block) • Rare cases of pancreatitis have been reported. Should be discontinued in the presence of persistent severe abdominal pain sometimes radiating to the back and which may or may not be accompanied by vomiting • Caution for use with ESRD, GI problems, hepatic liver disease, diabetic retinopathy

<p>GLP-1 Receptor Agonist (injectable)</p> <p><u>Liraglutide (Victoza)</u> <i>Initial Dose:</i> 0.6 mg SC daily <i>Average Dose (Max dose):</i> 1.2-1.8 mg SC daily (1.8 mg) <i>Expected HbA1C reduction:</i> 0.6-1.4% ODB Coverage: No</p> <p><u>Dulaglutide (Trulicity)</u> <i>Initial Dose:</i> 0.75mg (0.5 mL) SC qweekly <i>Average Dose (Max dose):</i> 1.5mg (1.5mg) <i>Special Instructions:</i> - Can be taken with or without meals <i>Onset/peak/duration:</i>24 hrs/48hrs/120hrs <i>Expected HbA1C reduction:</i> 0.6-1.4% ODB Coverage: No</p> <p><u>Semaglutide (Ozempic)</u> Initial Dose: 0.25mg SC weekly X 4 weeks (not a therapeutic dose) to be increased to 0.5mg SC weekly Average Dose (Max dose): 0.5mg (2mg) Special Instructions: - Can be taken with or without meals <i>Onset/peak/duration:</i>24 hrs/24-72hrs/1 week <i>Expected HbA1C reduction:</i> 0.6-1.4% (2.2% on 2mg) ODB Coverage: Yes</p>	<ul style="list-style-type: none"> • Liraglutide to be increased to ideal therapeutic dose of 1.2mg OD after 1 week as tolerated. Can further increase to 1.8mg OD if needed based on response after 1 week at 1.2mg • Dulaglutide to be increased to ideal therapeutic dose of 1.5mg q 1week after 1 week as tolerated. • Semaglutide to be increased to ideal therapeutic dose of 0.5mg in 4 weeks. Can increase further to 1mg in another 4 weeks if tolerating well. If additional glycemic control is needed after 4 weeks and tolerating well, the dose may be increased to 2mg once weekly. • Semaglutide's maximum recommended dose is 2mg once weekly which is included in the product monograph. Pre-filled pen delivering doses of 2mg per injection is not available in Canada at this time. The current method of administration for 2mg once weekly is two consecutive injections of 1mg. 	<ul style="list-style-type: none"> • Common adverse effects are nausea (10.7-18.6%), diarrhea (8.3-14.9%), headache (5.4-12.4%), vomiting (5.4-7.4%) and dyspepsia (2.1-7.0%) • Side effects usually improve over time • Should be stored in the refrigerator and unused medication discarded after 30 days • Increase in heart rate/ PR interval prolongation. Caution should be observed in patients with pre-existing conduction system abnormalities (first/second/third degree AV block) • Liraglutide is only approved for use with Metformin and/or a sulfonylurea in Canada • Semaglutide has been studied with Metformin, Metformin +Sulfonylurea, Metformin + SGLT2i, Metformin + Basal insulin • Dulaglutide can be used with Metformin, with Metformin & a Sulfonylurea, with mealtime insulin. • Dulaglutide has not been studied in combination with a basal insulin • Rare cases of pancreatitis have been reported. Should be discontinued in the presence of persistent severe abdominal pain and vomiting • Contraindicated with type 1 diabetes, DKA, personal or family history of medullary thyroid carcinoma or in patients with Multiple Endocrine Neoplasia syndrome • May slow absorption of medications; caution with medications that require rapid absorption (acetaminophen, pain medications) • Increased risk of hypoglycemia if used with sulfonylurea (assess need to decrease sulfonylurea dose by 50%) • In clinical trials, insulin dose was decreased by 20% at onset of Semaglutide treatment • Assess renal function prior to treatment and periodically thereafter • Semaglutide and Exanatide not recommended with eGFR <30, Liraglutide and Dulaglutide not recommended with eGFR <15 mL/min • Caution in patients with recent MI, unstable angina, CHF, IBS or gastroparesis (no studies) • Dulaglutide's day of weekly administration can be changed, if necessary, as long as the last dose was at least 3 days before • Semaglutide's day of weekly administration can be changed, if necessary, as long as the time between two doses is at least 2 days • Caution for use with ESRD, GI problems, hepatic liver disease, diabetic retinopathy
-------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------	---------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------	------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------

<p>SGLT2 Inhibitors</p> <p><u>Canagliflozin (Invokana)</u> <i>Initial Dose:</i> 100 mg daily qam ideally before meal <i>Average Dose (Max dose):</i> 300mg daily (300mg) <i>Onset/peak/duration:</i> Rapidly absorbed/1-2 hrs/approx. 24 hrs <i>Expected HbA1C reduction:</i> 0.5-0.7% * up to 2.56% with HbA1C >10% ODB Coverage: Yes</p> <p><u>Dapagliflozin (Forxiga)</u> <i>Initial Dose:</i> 5 mg daily with/without food <i>Average Dose (Max dose):</i> 10 mg daily (10 mg) <i>Onset/peak/duration:</i> Rapidly absorbed/2h/approx. 24 hrs <i>Expected HbA1C reduction:</i> 0.5-0.7% * up to 2.04% with HbA1C >9% ODB Coverage: No</p> <p><u>Empagliflozin (Jardiance)</u> <i>Initial Dose:</i> 10 mg daily with/without food <i>Average Dose (Max dose):</i> 25 mg daily (25 mg) <i>Onset/peak/duration:</i> Rapidly absorbed/1.5h/approx. 24 hrs <i>Expected HbA1C reduction:</i> 0.5-0.7% * up to 2.04% with HbA1C >9% ODB Coverage: Yes</p>	<ul style="list-style-type: none"> • Start Canagliflozin at 100mg and increase to 300mg if well tolerated and eGFR >60 mL/min • Start Dapagliflozin at 5mg od and increase to 10mg od if well tolerated and eGFR >60 mL/min • Start Empagliflozin at 10mg od and increase to 25mg od if well tolerated and eGFR >45mL/min • Canagliflozin should be discontinued when eGFR is <45 mL/min as it would not be effective in these patients and adverse reactions are more severe • Dapagliflozin should be discontinued when eGFR is <60 mL/min • Empagliflozin should be used with caution if eGFR 30-44 mL/min, discontinued when eGFR is <30 mL/min • SGLT2 inhibitors should be put on hold in patients with severe dehydration (i.e., vomiting and unable to keep down fluids) 	<ul style="list-style-type: none"> • Indicated as monotherapy in patients with type 2 diabetes for whom Metformin is inappropriate due to contraindications or intolerance • Indicated in combination therapy with Metformin, sulfonylureas or insulin (with or without Metformin) • Invokana and Forxiga are indicated in combination with Januvia • Common adverse effects are increased serum potassium >5.4 mEq/ml (12-27%) and >6.5 mEq/ml (2%), genital mycotic infections (7-11% in women and 3-4% in men), urinary tract infections (4-6%), nasopharyngitis (6-7%), polyuria (3-5%) • SGLT2 inhibitors can be added for glycemic control if eGFR >45 mL/min • SGLT2 inhibitors may be used for cardiorenal benefits in those with clinical CVD, A1C above target and eGFR >30 mL/min • Renal function should be assessed prior to initiation of and regularly after with more frequent monitoring for patients taking Canagliflozin or Empagliflozin with eGFR 45-60 mL/min • Monitor serum potassium levels periodically after initiating in patients with impaired renal function and in patients predisposed to hyperkalemia due to medications or other medical conditions • May increase the risk for ketoacidosis. Patients experiencing signs and symptoms of ketoacidosis (e.g., difficulty breathing, nausea, vomiting, abdominal pain, confusion, unusual fatigue or sleepiness) should be evaluated and SGLT2 inhibitor should be discontinued if acidosis is confirmed • LDL levels should be monitored due to dose dependent increases in LDL-C seen with therapy • Dapagliflozin should not be used in patients with active bladder cancer and should be used with caution in patients with a prior history of bladder cancer • Dapagliflozin is not recommended in combination with pioglitazone (Actos) • Dapagliflozin and Canagliflozin tablets contain lactose • May cause symptomatic hypotension due to intravascular volume depletion especially in patients with renal impairment (eGFR <60 mL/min), elderly, patients on other antihypertensives, or those with low systolic blood pressure. Assess volume status prior to initiation in patients at risk of hypotension and correct if
------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------	-------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------	----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------

		<p>depleted; monitor for signs and symptoms of hypotension after initiation</p> <ul style="list-style-type: none">• Not recommended for use with patients on loop diuretics• Can be used in mild-moderate hepatic impairment• Elderly patients (≥ 65 years) may have an increased risk of symptoms related to intravascular volume depletion (e.g., hypotension, orthostatic hypotension, dizziness, syncope, and dehydration) during therapy, especially with higher doses; elderly patients ≥ 75 years may experience a more pronounced risk. HbA1c reductions may be less in patients > 65 years compared to younger patients.• Patients should be educated about the increased risk for genital mycotic infections and/or urinary tract infections• Patients should be advised about the possible side effect of increased urination and encouraged to drink sugar-free liquids during the day to avoid dehydration• Increased risk for hypoglycemia if combined with sulfonylurea or insulin (may need to adjust diabetes medications)• A low-carbohydrate diet is not contraindicated for patients taking an SGLT2 inhibitor. However, there is increased risk for DKA while following a low-carbohydrate diet and taking an SGLT2i. Patients should be educated about this increased risk, signs and symptoms of DKA, and when to visit the Emergency Department
--	--	---------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------

<p>Combination Medications</p> <p><u>Janumet (Januvia and Metformin)</u> <i>Initial Dose:</i> 50/500mg bid <i>Average Dose (Max dose):</i> 50/1000mg bid (50/1000mg bid) <i>Special Instructions:</i> Take with meals to reduce GI side effects ODB Coverage: Yes</p> <p><u>Janumet XR (Januvia and Glumetza)</u> <i>Initial Dose:</i> 50/1000mg daily ideally with dinner <i>Average Dose (Max dose):</i> 50/2000mg od (50/2000mg od) <i>Special Instructions:</i> Take with meal to reduce GI side effects ODB Coverage: Yes</p> <p><u>Jentadueto (Trajenta and Metformin)</u> <i>Initial Dose:</i> 2.5/500mg bid <i>Average Dose (Max dose):</i> 2.5/1000mg bid (2.5/1000mg bid) <i>Special Instructions:</i> Take with meals to reduce GI side effects ODB Coverage: Yes</p> <p><u>Komboglyze (Onglyza and Metformin)</u> <i>Initial Dose:</i> 2.5/500mg bid <i>Average Dose (Max dose):</i> 2.5/1000mg bid (2.5/1000mg bid) <i>Special Instructions:</i> Take with meals to reduce GI side effects ODB Coverage: Yes</p>	<ul style="list-style-type: none"> • See the indications for medications included in combination tablet • Initiate at higher doses (not initial dose) if already taking Metformin at higher doses and tolerating well 	<ul style="list-style-type: none"> • See contraindications/precautions for medications included in combination tablet
--------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------	-----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------	--------------------------------------------------------------------------------------------------------------------------------------

<p><u>Kazano (Nesina and Metformin)</u> <i>Initial Dose:</i> 12.5/500mg bid <i>Average Dose (Max dose):</i> 12.5/1000mg bid (12.5/1000mg bid) <i>Special Instructions:</i> Take with meals to reduce GI side effects ODB Coverage: No</p> <p><u>Xigduo (Forxiga and Metformin)</u> <i>Initial Dose:</i> 5/850mg bid <i>Average Dose (Max dose):</i> 5/1000mg bid (5/1000mg bid) <i>Special Instructions:</i> Take with meals to reduce GI side effects ODB Coverage: No</p> <p><u>Invokamet (Invokana and Metformin)</u> <i>Initial Dose:</i> 50/500mg bid <i>Average Dose (Max dose):</i> 150/1000mg bid (150/1000mg bid) <i>Special Instructions:</i> Take with meals to reduce GI side effects ODB Coverage: No</p> <p><u>Synjardy (Jardiance and Metformin)</u> <i>Initial Dose:</i> 5/500mg bid <i>Average Dose (Max dose):</i> 12.5/1000mg bid (12.5/1000mg bid) <i>Special Instructions:</i> Take with meals to reduce GI side effects ODB Coverage: No</p> <p><u>Xultophy (Degludec and Liraglutide)</u> <i>Initial Dose:</i> 10u SC daily (pts naïve to basal insulin or GLP1 receptor agonist) OR</p>		
---------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------	--	--

<p>16u SC daily (pts currently using basal insulin or GLP1 receptor agonist) <i>Average Dose (Max dose):</i> N/A (50u daily) <i>Special Instructions:</i> -Take once daily at the same time with or without meals -Dose should be titrated up or down by 2u q3-4days to achieve fasting blood glucose targets -Basal insulin and/or GLP1 Receptor Agonist should be discontinued prior to starting Xultophy</p> <p><u>Soliqua (Glargine and Lixisenatide)</u> <i>Initial Dose:</i> 15u SC daily (pts taking <30u basal insulin daily) OR 30u SC daily (pts taking 30-60u basal insulin daily) <i>Average Dose (Max dose):</i> N/A(60u) <i>Special Instructions:</i> -Take once daily within an hour prior to the first meal of the day -Dose should be titrate dup or down by 2-4u every week to achieve fasting blood glucose targets -Basal insulin and/or GLP1 receptor agonist should be discontinued prior to starting Soliqua</p> <ul style="list-style-type: none"> • See dosing information for medications included in combination tablet for more information 		
------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------	--	--

Table 2 Notes:

- The implementer will adhere to the indications/contraindications outlined in this table
- The primary care provider or endocrinologist must be available to provide consultation as required
- The implementer will recommend hold of medication and contact the physician/nurse practitioner immediately if suspicion of a hypersensitivity reaction i.e. anaphylaxis, hives, rash etc.
- Implementer should follow up with the patient regarding medications discontinued or placed on hold in collaboration with physician/nurse practitioner.
- Women with type 2 diabetes who are planning a pregnancy should switch from noninsulin antihyperglycemic agents to insulin for glycemic control. Women with pregestational diabetes who also have PCOS may continue metformin for ovulation induction.
- Metformin and glyburide may be used during breastfeeding.

Table 3: Indications and Contraindications for Providing Insulin Samples

	Indications	Contraindications
Providing Insulin Samples	<ul style="list-style-type: none"> • The patient has obtained an Rx from their primary health care provider (PHCP) for the insulin to be provided or the RD or RN has been given a verbal or written order from the PHCP • Samples will be provided (as able) when needed for the timely initiation of insulin or due to the financial constraints of the patient. • The RN or RD will attempt to link the patient to any relevant financial assistance programs available for insulin and other diabetes management supplies i.e. trillium drug plan etc. • The insulin samples will be kept in a fridge with temperatures ranging from 2 – 8 degrees Celsius. The temp. will be monitored and recorded twice per day • Insulin samples should be inspected by the RD or RN prior to providing them to a patient to check expiration date and clarity of insulin (see contraindications) • The RN or RD will document expiry date, & lot # of the insulin dispensed 	<ul style="list-style-type: none"> • Insulin storage fridge temp. has dropped below freezing (insulin will need to be discarded) • Insulin has expired • Insulin has clumps, solid white particles or clear insulin appears cloudy

Table 3 Notes: Insulin samples cannot be donated by patients.

Appendix D

Implementer Competency Checklist

Implementer Name: _____

CDA Guidelines Chapter Reviews		
Diabetes Canada 2018 Clinical Practice Guidelines for the Prevention and Management of Diabetes in Canada	Date Reviewed	Signature
Pharmacologic Management of Type 2 Diabetes (including updated resource for 2020)		
Diabetes and Driving		
Appendix 9: Examples of Insulin Initiation and Titration Regimes in People with Type 2 Diabetes		
Appendix 5: Self-Monitoring of Blood Glucose (SMBG) Recommendation Tool for Healthcare Providers		
Appendix 7: Therapeutic Consideration for renal impairment		
Appendix 8: Sick Day Medication List		
CDA Building Competency in Diabetes Education: The Essentials	Date Reviewed	Signature
Chapter 6 - Treatment Modalities: Pharmacological Therapy		
Chapter 12 – Intensive Insulin Therapy		
Diabetes Canada Building Competency in Diabetes Education: Advancing Practice	Date Reviewed	Signature
Chapter 2 – Advanced Insulin Therapy		
Chapter 3 - Acute Care Issues: Driving Employment and Insurance Issues		
Canadian Injection Recommendations: FIT, 2020		

Performance Criteria	Observed ex. through mentoring or chart audits (Date)	Not Observed ex. discussion of cases (Date)	Comments
1. Able to identify the action, dosing, indications & contraindications, possible side effects of all antihyperglycemic oral medications identified in this directive			
2. Able to identify the pharmacokinetics and action times of all insulins described in this medical directive including onset, peak, duration			
3. Identifies potential side effects of insulin therapy and how to avoid/minimize and manage them (i.e., hypoglycemia, lipohypertrophy, weight gain, in rare cases allergy)			
4. Able to counsel Patient's on private and commercial driving recommendations when there is a higher risk for hypoglycemia i.e., taking insulin and/or sulfonylurea			
5. Identifies lifestyle factors, medications and/or medical conditions that can impact patient's glycemic control			
6. Describes basic physiologic insulin requirements in type 1 and type 2 diabetes in adults as well as usual starting doses based on age, weight, diagnosis, etc.			

Performance Criteria	Observed ex. through mentoring or chart audits (Date)	Not Observed ex. discussion of cases (Date)	Comments
7. Completes and documents comprehensive assessment of learning needs & provides timely, patient-centered education on diabetes management including insulin and medications			
8. Calculates, uses and evaluates insulin: carbohydrate ratios			
9. Calculates, uses and evaluates insulin sensitivity factor and correction doses			
10. Describes the purposes of consistent CHO intake and/or CHO counting and identifies potential advantages/disadvantages of each, according to patient's situation			
11. Identifies effect of alcohol consumption on blood glucose values and provides education and advice to minimize risk and prevent hypoglycemia			
12. Identifies dietary and/or insulin recommendations for physical activity.			
13. Identifies patterns of hyperglycemia or hypoglycemia or changes in routine that require adjustment of insulin, medications and/or other components of treatment plan			

Performance Criteria	Observed ex. through mentoring or chart audits (Date)	Not Observed ex. discussion of cases (Date)	Comments
14. Describes and/or has provided sick day management recommendations including dietary, medication/insulin adjustments as necessary			
15. Describes and/or has provided education and guidance for travelling with diabetes, including medication/insulin adjustment as needed			
16. Builds relationships with patients to promote self-care and learning; does not encourage ongoing dependence on health professionals for insulin adjustment i.e., increased patient confidence to self-adjust insulin			
17. Able to counsel patient around glucose meter usage and checking for accuracy i.e., lab/meter check or control solution			
18. Able to document process according to established standards and consult referring primary care provider for non-standard situations and/or failure of insulin dose adjustment to improve control.			
19. Familiarity with insulin pens and injections technique based on Canadian Recommendations FIT.			

Endocrinologist and Implementer Signature Sheet

I _____ have supervised _____
(Name of endocrinologist) (Name of Implementer)

with at least three patient cases and feel that they have achieved competency to adjust insulin and/or oral antihyperglycemic medications for patients with diabetes according to medical directive TCFHT-MD05 "Diabetes Management in Adults".

Signed _____ Endocrinologist

Signed _____ DEP Coordinator and/or Preceptor

Signed _____ Implementer

Date _____