There are a number of situations that patients taking insulin should be aware of how to troubleshoot.

Sick day management⁷²

- Counsel patients as follows when they are sick (e.g., vomiting, diarrhea, fever, dehydration):
 - Monitor blood glucose more often (e.g., every 1-2 hours)
 - If at risk of dehydration (i.e., cannot keep food/liquid down), temporarily stop agents from the SADMANS list (see <u>Type 2 diabetes and sick days: Medications to pause</u>)
 - Secretagogues (e.g., gliclazide, glyburide, repaglinide)
- **D**iuretics, direct renin
- inhibitors
 - Metformin

- ACE inhibitors
- Adjust insulin amount depending on changes in food/liquid intake
- · When feeling better (i.e., able to eat/drink well), restart agents from the SADMANS list and regular insulin routine

Diabetic Retuacidusis (DRA) and hyperusinular hypergryceniic state (ini	and hyperosmolar hyperglycemic stat	≥ (HHS)
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Risk factors³

- Type 1 diabetes (4.6-8 / 1,000 patient-years), type 2 diabetes (0.32-2 / 1,000 patient-years)^{73,74}
- New diabetes diagnosis
- Non-adherence to insulin therapy
- Thyrotoxicosis

· Severe infection

Myocardial infarction

Surgery

Trauma

Stroke

- Use of specific medications and drugs (e.g., SGLT2i, cocaine, atypical antipsychotics, interferon, diuretics, glucocorticoids, lithium)
- · Ultra low carb or keto diet

ARBs

NSAIDS

SGI T2is

- Pregnant patients³
 - Pregnant patients in DKA typically present with lower blood glucose levels than non-pregnant patients
 - There are case reports of euglycemic DKA in pregnancy
- Clinical presentation of DKA or HHS:

	Symptoms	Signs
DKA	Urge to breathe deeply, nausea, vomiting, abdominal pain, altered sensorium	Kussmaul respiration, acetone-odoured (fruity-smelling) breath
ннѕ	Polyuria, polydipsia, weakness	Extracellular fluid volume depletion, seizures, stroke-like state

Management of mild DKA at home (mild = alert; not drowsy, not in stupor or coma)³

- In usual care, patients can self-manage early DKA if clinically stable and able to drink fluids
- Monitor every 1-2 hours: blood glucose, nausea, vomiting, extreme thirst, ketones (with a reader/meter that accepts ketone strips, e.g., FreeStyle Libre[®], FreeStyle Insulinx[®], FreeStyle Precision Neo[®])
- Hospitalization³
 - For patients with clinical decline (and/or a positive ketone test), recommend an evaluation at the emergency department given the concern of a rapid clinical decline (patients should not drive themselves)

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Physical activity

- Encourage patients to participate in regular physicial activity
- Insulin dose adjustments may be needed due to physicial activity
 - Exact insulin dose adjustments cannot be provided
- Adjustments will range (e.g., minimal to 50%), depending on the type/duration of exercise and blood glucose levels
- Counsel patients who plan to participate in physical activity:75,76
 - Monitor blood glucose before, during and a few hours after any physical activity
 - Adjust insulin as needed
 - Always carry rapid-acting carbohydrates (e.g., dextrose tablets)
 - · Consider insulin timing when physical activity is planned
- Avoid injecting insulin into subcutaneous tissue next to the primary muscle used (activity will increase insulin absorption)
- Eat before before any physical activity
- Ensure proper hydration and watch for signs and symptoms of dehydration (e.g., increased thirst, nausea, severe fatigue, blurred vision or headache)³

- Driving
- Risk
 - · Diabetes, the use of insulin and its complications can affect driving performance
 - Possible risks include: impaired sensory or motor function, impaired cognition, diabetic eye disease, nerve damage, kidney disease, cardiovascular disease, peripheral vascular disease, stroke and incidents of hypoglycemia³
- Assessment
 - All drivers taking insulin should have fitness to drive assessed every 2 years³
 - Driver's license may be suspended if a patient is determined unfit to drive or experiences an accident caused by hypoglycemia⁷⁸
 - Medical professionals in Ontario have a legal obligation to report to the appropriate regulatory body patients who have conditions
 that impair their driving ability (see <u>Reporting a driver for medical review</u>)^{3,77}
 - Counsel patients on insulin secretagogues and/or insulin to (see Drive safe with diabetes").³
 - Measure blood glucose levels immediately before driving (must be >4.0 mmol/L)
 - · Re-test at least every 4 hours while driving
 - At least every 2 hours if the patient has a history of severe hypoglycemic episodes (e.g., loss of consciousness) or hypoglycemia unawareness (e.g., lack of early warning symptoms of hypoglycemia, such as tremor, sweatiness and palpitations)
 - If blood glucose <4.0 mmol/L, treat hypoglycemia (i.e., ingestion of 15g of carbohydrate). Re-test and wait 40 minutes after blood glucose returns to >4.0 mmol/L before driving
 - Eat next meal or snack (containing carbohydrate and protein) within an hour
 - Always keep an emergency supply of fast-acting carbohydrates (e.g., dextrose tablets) and a glucose monitoring system inside the vehicle Commercial vehicle licensing^{3,78}
 - Canadians with diabetes (with or without insulin) can be licensed to drive a commercial vehicle in Canada (and now the United States) if medical standards are met

Lipohypertrophy⁵¹

- Presentation
 - Lipohypertrophic area(s) can develop under the skin where the same injection or infusion site is used repeatedly
 - · Lipohypertrophy usually presents as thickened or rubbery lesions (can vary in size and shape)
 - Some lesions are easily seen (e.g., a large bulge), while others require detected by palpation (e.g., a hard lump) or ultrasound
- Risk factors
 - Lipohypertrophy can develop from repeated use of the same area, reusing insulin needles, more frequent insulin injections, higher dose insulin and a lack of systematic rotation
- Effect on insulin
 - Lipohypertrophy can decrease the rate of insulin absorption or make it more erratic/delayed, resulting in higher doses of insulin needed to achieve glycemic targets
- Prevention
 - · Assess patients on insulin regulary for lipohypertrophy
 - Palpate injection area(s) regularly during patient appointments
 - Counsel patients on proper insulin injection techniques and pen needle use (see <u>Type 2 diabetes: insulin therapy > Safe insulin</u> injection techniques and pen needle use)
- Treatment
 - Have patient avoid injecting in the site
 - Lipohypertrophy may take 3-6 months to reduced or resolve

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Insulin and pre-surgery/procedure⁷⁹

- Counsel patients
 - Night before surgery:
 - Consider reducing long-acting, intermediate-acting or premixed insulin by 20-25%
 - Stop prandial (bolus) insulin when fasting begins
 - Morning of surgery (defer to any pre-op information patient has been given if available):
 - Patients on a twice-daily dosing of basal insulin should consider reducing the normal morning dose by 20%
 - If morning blood glucose > 6.7 mmol/L: reduce intermediate-acting or premixed insulin to 50% of the usual dose
 - If morning blood glucose < 6.7 mmol/L: do not take intermediate-acting or premixed insulin. This will minimize the risk of hypoglycemia



Travel⁸⁰⁻⁸²

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- The primary goal is to avoid/prevent hypoglycemia during travel
- Counsel patients on general diabetes travel tips (see <u>Air travel: Traveling can be a breeze if you follow a few tips</u>ⁱⁱⁱ)
- Recommendations for insulin adjustment during air travel vary significantly and are mostly based on expert opinion

Travel	Insulin adjustment	Example
Eastward travel, crossing more than 5 time zones (lose hours, shorter day) • Example: Toronto, Ontario to London, England	 Once-daily insulin: Inject the usual dose on the day of travel Inject 2/3 of the usual dose on the day destination is reached (travellers lose a few hours and may consume fewer calories) Patients on long-acting insulin can also keep time of administration on Canadian time 	 Glargine 60 units at bedtime on day of travel in Canada Glargine 40 units at bedtime in Europe on day 1, then 60 units every night at bedtime thereafter
	 Twice-daily insulin: Inject 2/3 of the usual morning dose on the day of arrival at the destination Usual dinner dose is the same Usual dose followed from day 2 onwards 	 Detemir 30 units twice daily in Toronto In Europe, detemir 20 units in the morning and 30 units in the evening, then 30 units twice daily thereafter
	 Basal-prandial (bolus) therapy Usual dosing schedule If a meal is missed during travel, then the prandial (bolus) dose should be skipped 	 Glargine 60 units at bedtime and lispro 20 units three times daily with meals – no change
 Westward travel, crossing 5 or more time zones (gain hours, longer day) Example: London, England to Toronto, Ontario 	 No insulin dose change, but may require extra prandial (bolus) insulin with extra meal if prandial (bolus) insulin is used 	No insulin dose change
Travel within North America	No insulin dose change	No insulin dose change

Specific diets

- Refer patients with specific diet requirements to a dietitian (see Local services for patients living with type 2 diabetes^{IV})
- For information on common specific diets for patients with type 2 diabetes:
- Patients observing Ramadan⁸³
- Low carbohydrate diets⁸⁴
- Intermittent fasting⁸⁵



Glucocorticoid-induced hyperglycemia^{86,87}

- Glucocorticoid-induced hyperglycemia can occur in patients with or without diabetes
- Monitoring
 - Monitor blood sugar three times daily
- Indications to consider treatment
 - Blood glucose >12 mmol/L on two occasions within a 24-hour period (consider checking ketones if indicated)
- Treatment options
 - If NOT on insulin: adding a sulfonylurea or adjusting current sulfonylurea dose may be considered. Insulin may be needed.
 - If ON insulin: adjust basal insulin daily (may need >30% increase)
- Treatment considerations should be individualized based on glucocorticoid drug (e.g., prednisone has mid-day peak, dexamethasone is longer acting), treatment duration, and patient risks of hyperglycemia and hypoglycemia. A second opinion from a specialist may be considered (e.g., e-Consult)
- Careful monitoring and subsequent decreases in antihyperglycemic therapy is prudent when steroid dose is tapered and/or when clinical status changes

Patient resources

- [i] RxFiles Type 2 diabetes and sick days: Medications to pause
- [ii] Diabetes Canada Drive safe with diabetes
- [iii] Diabetes Canada Air travel: Traveling can be a breeze if you follow a few tips
- [iv] Centre for Effective Practice local services for patients living with type 2 diabetes

References

See Type 2 Diabetes: Insulin Therapy tool

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The Complex insulin events tool ('Tool') was developed as part of the Knowledge Translation in Primary Care Initiative, led by the Centre for Effective Practice, in collaboration with the and Nurse Practitioners' Association of Ontario. Clinical leadership for the development of the Tool was provided by Dr. Risa Bordman and was subject to external review by health care providers and other relevant stakeholders. This Tool was funded by the Government of Ontario as part of the Knowledge Translation in Primary Care Initiative.

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