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#### BCCH DIABETIC KETOACIDOSIS PROTOCOL TOOLKIT

The BCCH DKA Protocol Toolkit contains the following documents:

- the BCCH DKA Protocol Toolkit cover document (2022/04/03 version)
- the BCCH DKA Medical Protocol (2020/02/08 version) PLAIN PDF FORMAT\*
- the BCCH DKA Nursing Protocol (2022/04/03 version)
- the BCCH DKA Flowsheet (2020/12/13 version)
- the BCCH DKA Sample Prescriber Order Sheet (2019/10/08 version)
- the BCCH DKA Recipes for Making Solutions handout (2019/10/08 version)
- the BCCH DKA Glucose, Insulin and Fluid Management handout (2022/04/03 version)

Each of these documents is also available individually for download from our website: www.bcchildrens.ca/health-professionals/clinical-resources/endocrinology-diabetes/dka-protocol

\*The BCCH DKA Medical Protocol can also be downloaded in **fillable PDF format** from our website: www.bcchildrens.ca/endocrinology-diabetes-site/documents/dkaprtfill.pdf

Following the publication of the PEKARN DKA FLUID Trial [New Engl J Med 2018;378(24):2275-2287], which demonstrated the safety of more-aggressive fluid replacement regimens than are used in current DKA protocols, the Division of Pediatric Endocrinology & Diabetes has updated the BCCH DKA Protocol. Our 2019 revision aligns closely with the protocol developed by TREKK (Translating Emergency Knowledge for Kids), which is designed for the initial management of pediatric DKA in most Canadian emergency departments, as well as with the DKA algorithm developed by the Canadian Pediatric Endocrine Group, which is designed for ongoing inpatient management of DKA. The 2019 revision is also aligned with the Clinical Practice Consensus Guidelines 2018 of the International Society for Pediatric and Adolescent Diabetes (ISPAD).

The major modifications from the previous version (dated 2015/10/07) of the protocol include:

- more-aggressive fluid boluses are suggested at the start of therapy: all patients with DKA should receive a 10-mL/kg bolus of normal saline at the beginning, and the majority will receive a second 10-mL/kg bolus to follow
- fluid infusion rate calculations have been simplified
- fluid rehydration rates will reflect a goal to correct losses over a 36-h period (previously this was 48 h)

We hope that you will find these materials to be helpful in managing pediatric cases of diabetic ketoacidosis. Please do not hesitate to contact the Endocrinology & Diabetes Unit at BCCH for any help in implementing this protocol at your health-care centre in British Columbia. We also welcome any suggestions to make this material more useful to your practice.

## **FIRST 60 MIN**

# TIME = 60 MIN-36 HOURS

#### BC CHILDREN'S HOSPITAL DIABETIC KETOACIDOSIS PROTOCOLA

#### FOR CHILDREN AGES 1 MONTH TO 19 YEARS

THIS PROTOCOL IS ALSO AVAILABLE IN FILLABLE PDF FORMAT



### 0. ABCs, vital signs (with BP), neurovitals signs. Place large-bore IV. Draw labs. Confirm DKA: plasma glucose (PG) >11 mmol/L, moderate—large ketonuria or β-hydroxybutyrate ≥3.0 mmol/L, and venous pH <7.3 or serum HCO<sub>3</sub><sup>-</sup> <15 mmol/L.<sup>c</sup> Consider possibility of an element of hyperglycemic hyperosmolar state.<sup>B</sup>

- 1. Measure body weight (BW) in kilograms ...... kg
- 2. Give 0.9% saline (normal saline, NS) resuscitation bolus<sup>D</sup>
  - recommended amount: 10 mL/kg BW over 30 minutes.....(2) \_\_\_\_\_ mL
- 3. Repeat with second bolus of NS if persistent tachycardia, prolonged cap refill (>2 sec), cool extremities:
  - recommended amount: 10 mL/kg BW over 30 minutes......(3) mL
- 4. Begin rehydration, calculated for even correction over 36 hours, based on admission BW:<sup>E</sup>
  - 5–10 kg BW: 6.5 mL/kg/h
  - 10-20 kg BW: 6 mL/kg/h
  - 20-40 kg BW: 5 mL/kg/h
  - >40 kg BW: 4 mL/kg/h, maximum 250 mL/h ......(4) \_\_\_\_\_ mL/kg/h
- 5. Calculate **total** hourly fluid rate to be given for 36 hours: multiply (1) and (4).....(5) mL/h
- 6. Use NS with KCl 40 mEq/L (Bag A) as initial rehydration fluid, at rate determined in (5), ensuring that patient has voided and has plasma K<sup>+</sup> <5 mmol/L before adding potassium to the IV fluids.
- 7. At 60–120 minutes after starting the first fluid bolus, make up and start a piggyback insulin drip at 0.05–0.1 units/kg BW/h (Bag C):<sup>F</sup>
  - 50 units insulin regular (Humulin® R or Novolin® Toronto) in 500 mL NS or D10/NS
  - run at 0.5–1 mL/kg BW/h ......(7) \_\_\_\_\_ mL/h
- 9. Aim to keep PG ~8–12 mmol/L by titrating the rates of these two solutions, keeping the combined rate at (8)<sup>G</sup>. Continue this for the next 6–12 hours, monitoring as below.
- 10. At 4–6 hours after initial fluids and if corrected plasma Na<sup>+</sup> is ≥145 mmol/L, stable or increasing, switch Bag A to 0.45% saline w/ 40 mEq/L KCl and Bag B to D10–D12.5/0.45% saline w/ 40 mEq/L KCl at the rate as in (8)<sup>H</sup>.

#### Rationale & Notes:

APlease note that this protocol is designed as an algorithm for treating the majority of cases of DKA in infants, children and adolescents. It cannot replace careful clinical observation and judgment in treating this potentially very serious condition. If you have questions or problems related to the management of DKA or diabetes, please feel free to contact the BCCH Pediatric Endocrinologist on call.

<sup>B</sup>Hyperglycemic hyperosmolar state (HHS) should be suspected when there is significant hyperglycemia (>33 mmol/L) and hyper-osmolality (>330 mOsm/L) without ketosis or acidosis (bicarbonate >15 mmol/L, venous pH >7.3). A mixed picture of DKA and HHS is possible. Mild hyperglycemia, even with ketones and mild acidosis, can often be managed without IV fluids or IV insulin.

<sup>c</sup>Rapid, deep mouth-breathing (Kussmaul respiration) often dries out the oral mucosa, making the child appear more dehydrated than s/he really is. The hematocrit and other clinical signs (decreased capillary refill) are more accurate measures of dehydration.

<sup>D</sup>Recent research shows that most children with moderate—severe DKA will require a 20 mL/kg resuscitation fluid bolus to restore perfusion, prior to the rehydration phase.

<sup>E</sup>Recent research shows that DKA can be safely corrected over a 24- to 48-h period. This protocol is designed to correct a 10% fluid deficit (100 mL/kg) evenly over 36 h.

FIV insulin boluses are always contraindicated. Insulin given in the first 1–2 h of DKA repair is thought to increase mortality. This insulin rate fully inhibits ketogenesis and gluconeogenesis and should be maintained if possible. If unable to keep PG >8 mmol/L<sup>G</sup>, drop the insulin rate by 25–50%.

11. Re-evaluate appropriateness of replacement fluid type frequently, anticipating the need to add or increase Na<sup>+</sup>, K<sup>+</sup>, dextrose, etc.

• dextrose<sup>G</sup>: aim to keep the PG ~8–12 mmol/L range

• sodium<sup>H</sup>: corrected Na<sup>+</sup> <145 mmol/L, or falling regardless of level:

continue NS

corrected Na<sup>+</sup> ≥145, stable or increasing, switch to ½NS after 4–

6 h

• potassium<sup>I,J</sup>: patient urinating and K<sup>+</sup> remains <5: continue KCl 40 mmol/L

may give 50% of K<sup>+</sup> as acetate or phosphate

• bicarbonate<sup>K</sup>: NaHCO₃ is **not** generally recommended

- 12. Children with DKA have high risk for acute kidney injury (AKI). Use Schwartz formula to calculate expected baseline creatinine (EBC).<sup>L</sup>
- 13. Close neurological observation and frequent rousing of the child with finger-pokes to detect any changes consistent with cerebral edema. Follow Glasgow Coma Scale. Severe headache, change in sensorium or BP, dilated pupils, bradycardia, irregular breathing, posturing and incontinence are signs of impending deterioration. Rapid intervention is imperative:
  - airway / breathing / circulation
  - elevate head of bed
  - decrease all fluid bags to 5 mL/h pending physician reassessment
  - mannitol 20% (0.5–1 g/kg, 2.5–5 mL/kg IV over 15 min) or NaCl 3% (2.5–5 mL/kg
     IV over 15 min)<sup>M</sup>
  - consider intubation and mild hyperventilation (keep pCO<sub>2</sub> >22 mg Hg) for impending respiratory failure
  - arrange CT when stable

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- 14. Follow laboratory parameters (use of a flowsheet is highly recommended):
  - follow PG by meter every 30–60 min<sup>G</sup>: does child respond to the poke?
  - follow Na<sup>+</sup>, K<sup>+</sup>, Cl<sup>-</sup>, HCO<sub>3</sub><sup>-</sup>, anion gap, urea, creatinine, venous pH every 2–4 hours H, I, K; Ca<sup>2+</sup>, Mg<sup>2+</sup> and P<sub>i</sub> every 2–4 hours if giving phosphate<sup>J</sup>
  - ullet follow (preferably) plasma eta-hydroxybutyrate every 2–4 hours or urine ketones with each void
- 15. Re-evaluate appropriateness of replacement fluid type frequently, anticipating the need to increase or decrease Na<sup>+</sup>, K<sup>+</sup>, dextrose, etc.

<sup>G</sup>Keeping the PG in the ~8–12 mmol/L range allows for a buffer against hypoglycemia and a too-rapid fall in plasma osmolality<sup>H</sup>. The "two-bag method" (see our <u>DKA Nursing Protocol</u>) is a handy way to adjust the glucose without altering the Na<sup>+</sup> or K<sup>+</sup> delivery. It also allows for a faster response to PG changes, and it decreases nursing and pharmacy workload and costs.

<sup>H</sup>The introduction of hypotonic fluids must be considered carefully. The corrected Na $^+$  should be calculated and followed closely: corrected Na $^+$  = [measured Na $^+$  + 0.36×(PG–5.6)]. If corrected Na $^+$  falls or fails to rise as the PG falls, this could indicate excess free-water administration. It is also helpful to monitor the active osmolality [PG + 2×(Na $^+$  + K $^+$ )], which should not fall >0.5 mOsm/kg/h. If the corrected sodium is ≥145 mmol/L and stable and the active osmolality has been dropping slowly, switching to ½NS can be considered after 4–6 h of fluids. An elevated measured Na $^+$  in the face of hyperglycemia indicates severe dehydration and an element of the hyperglycemic hyperosmolar state. Such patients should be rehydrated using fluids with higher osmolar content (e.g. NS) for longer time periods (10–12 h).

'Serum K $^+$  levels are usually normal at diagnosis and fall precipitously with treatment. An IV fluid containing 20–40 mmol/L K $^+$  is usually required to keep the serum K $^+$ >3.0 mmol/L. Begin K $^+$  and insulin together. Oral/nasogastric KCl boluses (0.5–1 mmol/kg BW) may also be administered.

<sup>J</sup>While there is no proven benefit to using potassium phosphate or acetate, it does have the theoretical advantage of repleting the severe phosphate deficit of DKA and/or ameliorating the hyperchloremia which inevitably occurs during DKA treatment. If phosphate is given, serum calcium, magnesium and phosphate levels should be monitored closely.

 $^{K}$ The acidosis of DKA is due to both ketoacids and lactic acid, and these resolve with fluid and insulin replacement. There is no evidence that NaHCO<sub>3</sub> is either necessary or safe in DKA, but its use has a number of deleterious effects: paradoxical CNS acidosis, hypokalemia, hyperosmolality, delayed clearance of ketones, and cerebral edema. NaHCO<sub>3</sub> in DKA should only be considered if pH <6.9 or cardiac failure.

LEBC ( $\mu$ mol/L) = 36.5 × height (cm)/120. Measured creatinine 1.5–1.99× EBC = Stage 1, 2–2.99× EBC = Stage 2, ≥3× EBC = Stage 3 AKI.

 $^{M}\text{Subclinical brain swelling}$  is common in children with DKA. Cerebral edema (CE) accounts for more than half of the  $\sim\!1\text{--}5\%$  mortality rate of DKA in children. At highest risk are newly diagnosed patients, those aged <5 years, and those with initial pH <7.1 or pCO $_2$  <18. The exact etiology of CE remains unclear. Resuscitation is successful in only 50% of cases.

#### Accompanying documents on our website:

- DKA Flowsheet and DKA Sample Physician Order Sheet
- DKA Glucose, Fluid and Insulin Management
- <u>DKA Nursing Protocol</u> (including the "two-bag" method)
- DKA Recipes for Making Solutions

#### **BC CHILDREN'S HOSPITAL ENDOCRINOLOGY & DIABETES UNIT**

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#### BCCH DIABETIC KETOACIDOSIS NURSING PROTOCOL

Diabetic ketoacidosis (DKA) involves a combination of hyperglycemia, acidosis, and ketones. It is diagnosed when (1) the blood glucose is >11 mmol/L; (2) capillary pH is <7.3 and/or capillary bicarbonate is <15 mmol/L; and (3) ketones are present in the blood and/or urine (see below). It usually takes days to develop DKA, but it can take hours in children with acute illness, insulin omission, or insulin pump site problems.

#### Causes of DKA Include:

- undiagnosed type 1 diabetes
- insulin omission or manipulation
- inadequate insulin dosing and monitoring during periods of increased insulin needs:
   (illness, infection, major stress, puberty, pregnancy)
- insulin pump misuse or infusion site disconnection, kinking or failure

#### Signs and Symptoms of DKA Include:

- polyuria
- polydipsia
- dehydration
- weight loss
- lethargy
- nausea, vomiting and abdominal pain
- fruity or acetone-smelling breath
- flushed face
- confusion
- hyperventilation and Kussmaul breathing (rapid, deep, sighing mouth-breathing)
- $\land$  heart rate and  $\land$  respirations, and possibly  $\lor$  blood pressure

Acute dehydration must be treated with IV fluid replacement. Overhydration, correcting the hyperglycemia too quickly, the use of insulin in the first 1 to 2 hours of fluid therapy, and the use of bicarbonate have been implicated in causing cerebral edema in DKA, which can be fatal. Hydration should be cautious, according to the BCCH DKA Protocol.

#### NURSING MANAGEMENT OF DKA IN THE EMERGENCY ROOM

- 1. Patient should be kept NPO.
- 2. A measured weight is essential for rehydration calculations. The ER physician will estimate the degree of dehydration and the fluids required, using the BCCH DKA Protocol.
- 3. Baseline and then 1-hourly vital signs and neurovital signs.
- 4. Apply pulse oximetry and cardiac monitor;  $O_2$  via mask if saturation is low.
- 5. Measure blood glucose with hospital meter. If bloodwork is done at the same time, a drop from the lab sample may be used to do this. If the meter reads "HI", the blood glucose is ~30 mmol/L or greater, and the physician may request that labwork be drawn to obtain the actual blood glucose level.
- 6. Initial bloodwork: CBC, glucose, sodium, potassium, chloride, bicarbonate, osmolality, urea, creatinine, ionized calcium, phosphorus, venous blood gas and serum ketones/β-hydroxybutyrate (where available); and urine for ketones and glucose. Bloodwork can be done with the IV start. If IV start is difficult, call the Lab to do a stat finger sample rather than waiting for the IV line to be initiated.
- 7. Start one large-bore IV line. Three infusions will be Y'd into this line.
- 8. All patients will receive a 10 mL/kg normal saline (NS, sodium chloride 0.9%) bolus over 30 minutes once the IV is in. At the physician's discretion, moderately to severely dehydrated patients may receive a second 10 mL/kg NS bolus over 30 to 60 minutes. Very rarely, patients may require a third NS bolus.
- 9. After the initial NS fluid boluses, the desired fluid is NS + 40 mmol KCl/L, assuming serum potassium level is not elevated and the patient is urinating. The ER physician will calculate the rate of this from the BCCH DKA Protocol.
- 10. NOTE THAT INSULIN IS NOT GIVEN IN THE FIRST 1 to 2 HOURS OF DKA MANAGEMENT.
- 11. Set up the "two-bag system". This consists of two IV bags (A and B) with equal electrolyte concentration, one containing no dextrose, the other 10 or 12.5% dextrose. They are administered simultaneously. The concentration of dextrose is easily changed by adjusting the proportions of the two bags contributing to the total rate. The total rate is determined by the child's degree of dehydration, according to the BCCH DKA Protocol. The insulin infusion (Bag C) will eventually be Y'd into these bags (see below).

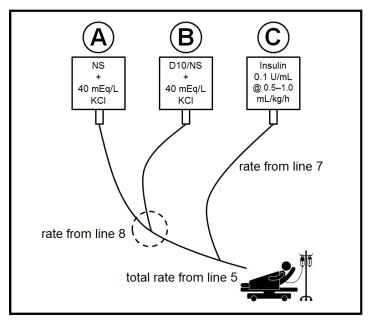
In the "two-bag system", **Bag A** is generally NS + 40 mEq/L KCl, and **Bag B** is usually D12.5/NS + 40 mEq/L KCl (or D10/NS + 40 mEq/L KCl if D12.5/NS is not available). The BCCH Pharmacy has prepared a "recipe book" for preparing these solutions from available IV solutions, which is available from the *Parenteral Drug Manual* on the C&W Intranet and on Endocrinology's website (see below). The following solutions should also be available pre-made in the Emergency Room and on T7 for after-hours use:

- NS + 40 mEq/L KCl
- D12.5/NS + 40 mEq/L KCl
- $\frac{1}{2}$ NS + 40 mEq/L KCl
- D12.5/ $\frac{1}{2}$ NS + 40 mEq/L KCl

#### EXAMPLE OF THE "TWO-BAG SYSTEM"

Example: IV rate from protocol line 8 = 100 mL/h (this does not include insulin infusion rate) rate Bag A + rate Bag B = 100 mL/h

rate Bag A	rate Bag B	final dextrose
no dextrose	D10	concentration
(mL/h)	(mL/h)	(%)
100	0	D0
80	20	D2
60	40	D4
40	60	D6
20	80	D8
0	100	D10



after: Grinberg A et al, Journal of Pediatrics 1999;134(3):376-378.

12. Insulin is started 1 to 2 hours after initial DKA fluid management is begun. To prepare the insulin infusion, 50 units (0.5 mL) of short-acting insulin regular (Humulin® R or Novolin® Toronto) is added to a 500-mL bag of NS (or to D10/NS, if ordered). This is a concentration of 0.1 units/mL. The insulin is drawn up in a tuberculin syringe with a  $1\frac{1}{2}$ -inch needle, so that the insulin is injected past the plastic port of the IV bag. **Do not use an insulin syringe**. Mix fluid continually while injecting, to prevent the insulin from settling in the port. Flush the tubing with 50 mL of the insulin solution to saturate insulin binding sites. **This preparation of the insulin bag requires a double-check if not performed in the Pharmacy**. The insulin infusion (**Bag** C) is Y'd into the lowest port on the IV, closest to the patient, and is usually run at 0.5 to 1 mL/kg/h (which is 0.05 to 0.1 units/kg/h). An

insulin bolus is never given. This high dose of insulin is required to reverse the ketosis. The BG level will fall quite rapidly in the first hour or two with the initial fluid management, even before insulin is started, secondary to improved renal clearance and hemodilution. Thereafter, one should aim for a fall in BG of  $\sim 3$  to 5 mmol/L/h. As the blood glucose approaches 15 mmol/L, or if it is dropping >5 mmol/L/h, the total rate of the insulin infusion will remain the same, but the rate of the no-dextrose **Bag A** will be decreased, and the rate of the high-dextrose **Bag B** will increased by the same amount. The target is to have the BG in the 8 to 12 mmol/L range, both to minimize glycosuria, and to allow for a buffer against hypoglycemia.

- 13. The insulin infusion is discontinued once the blood pH returns to normal and the patient is ready to switch to subcutaneous insulin. This is usually within 24 to 36 hours. Pharmacy prepares a new insulin infusion bag every 24 hours. The tubing is replaced every 96 hours, as per BCCH Nursing Policy and Procedure Manual CV.01.05 Administration Set Priming and Loading and Initiating or Changing the Infusion, available on the C&W ePOPS website.
- 14. Depending on the patient's progress, the solutions may eventually (e.g. after 6 to 12 hours) be changed to  $\frac{1}{2}NS + 40$  mEq/L KCl and D10-12.5/ $\frac{1}{2}NS + 40$  mEq/L KCl.

#### NURSING CARE

- blood glucose by meter and/or lab every 30 to 60 minutes
- electrolytes, lab glucose, blood ketones, venous gas every 2 to 4 hours as ordered
- record nursing results on DKA flowsheet
- vital and neurovital signs on admission and then hourly
- monitor for headache, abnormal respirations or behavioral changes
- NPO until rehydrated and glucose is stabilized
- ice chips may be allowed, at physician's discretion
- check urine for ketones with each void
- strict intake and output

#### CORRELATION OF BLOOD AND URINE KETONES

Urine	ketones	Blood ketones (β-hydroxybutyrate)		
negative	<0.5 mmol/L	≤0.5 mmol/L		
trace	0.5 mmol/L	0.6-0.9 mmol/L		
small	1.5 mmol/L	1.0-1.4 mmol/L		
moderate	4 mmol/L	1.5-2.4 mmol/L		
large	8 mmol/L	2.5-2.9 mmol/L		
very large	16 mmol/L	≥3.0 mmol/L		

#### MONITOR FOR CEREBRAL EDEMA

Cerebral edema occurs in  $\sim$ 0.5% of children presenting in DKA, and it has a mortality of  $\sim$ 25%. At highest risk are (1) children newly diagnosed with diabetes, (2) younger children, and (3) children with the greatest degree of dehydration and acidosis. Symptoms include:

- headache
- inappropriate lowering of heart rate
- recurrence of vomiting
- changes in neurological status (restlessness, irritability, drowsiness, incontinence, cranial nerve palsies, altered pupillary reactivity, etc.)
- rising blood pressure
- oxygen desaturation

If you suspect cerebral edema, notify the physician immediately. Elevate the head of the bed. Decrease all IV bags to 5 mL/h pending physician reassessment. Be prepared to call the code team, and ensure that IV mannitol (available on the BCCH wards) and/or 3% saline is ready at hand.

#### TRANSFER TO WARD

This may happen any time after the child is stabilized. This generally means that the patient's cardiovascular and CNS status is stable, and the patient has **Bags** A, B and C hanging. Disposable insulin pens with basal (Basaglar® and Levemir®), intermediate-acting (Humulin® N and Novolin® NPH) and rapid-acting (Humalog® and NovoRapid®) insulin will be available on the wards, in preparation for a switchover to SQ insulin. Nursing care continues as above until the insulin infusion is discontinued.

Subcutaneous insulin is started when acidosis is corrected and the child is ready to eat. The blood pH will be normal, and serum ketones (see below) will have normalized, but

ketones will likely still be present in the urine. For rapid-acting insulin (Humalog® or NovoRapid®), the injection is generally given immediately before breakfast or dinner, and the insulin infusion is turned off 20 to 30 minutes after the injection. An injection of basal insulin (Basaglar® or Levemir®) is generally given at the same time. The physician may choose to continue the IV fluids for another 12 to 24 hours to complete rehydration. Labwork will be discontinued once the child's pH and electrolytes have returned to normal.

For newly diagnosed children, diabetes education is initiated with the family as soon as possible. Children who are not newly diagnosed will need a reassessment of their diabetes management.

#### INTERNET LINKS

The following resources are all available from our DKA Protocol webpage:

- DKA Protocol Toolkit
- DKA Medical Protocol (PLAIN PDF FORMAT)
- DKA Medical Protocol (FILLABLE PDF FORMAT)
- DKA Nursing Protocol
- DKA Flowsheet
- DKA Sample Prescriber Order Sheet
- DKA Recipes for Making Solutions
- DKA Glucose, Insulin and Fluid Management
- Blood Glucose and Insulin Record for Conventional Insulin Regimens
- Blood Glucose and Insulin Record for MDI

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Wolfsdorf JI, Glaser N, Agus M, Fritsch M, Hanas R, Rewers A, Sperling MA, Codner E. ISPAD Clinical Practice Consensus Guidelines 2018: Diabetic ketoacidosis and the hyperglycemic hyperosmolar state. *Pediatric Diabetes* 2018:19(Suppl 27):155-177.

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Grimberg A, Cerri RW, Satin-Smith M, Cohen P. The "two bag system" for variable intravenous dextrose and fluid administration: Benefits in diabetic ketoacidosis management. *Journal of Pediatrics* 1999;134(3):376-378.

Translating Emergency Knowledge for Kids (TREKK Canada): www.trekk.ca.

BC Children's Hospital SHOP (Shared Health Organizations Portal): shop.healthcarebc.ca/phsa/bc-cnw-hospitals.



#### BCCH ENDOCRINOLOGY & DIABETES UNIT DIABETIC KETOACIDOSIS FLOWSHEET

DATE: TIME:						
HEART RATE						
RESPIRATORY RATE						
BLOOD PRESSU	RE					
GLASGOW COM	A SCALE					
NEURO ✓ DON	E?					
BLOOD	METER					
<b>G</b> LUCOSE	LAB					
URINE KETONES						
Nurse's Initi	IALS					
VENOUS PH						
BICARBONATE:	VENOUS					
HCO <sub>3</sub>	SERUM					
BASE DEFICIT						
SODIUM: NA+						
POTASSIUM: K	+					
CHLORIDE: CL						
ANION GAP: [NA+ + K+ - CL HCO3-]						
β-Hydroxybutyrate						
"CORRECTED" SODIUM: NA* + 0.36×[GLUCOSE-5.6]						
"ACTIVE" OSMOLALITY: GLUCOSE + 2×[Na++K+]						
UREA						
CREATININE						
CALCIUM						
PHOSPHATE						
PHYSICIAN'S INITIALS						

## PRESCRIBER'S ORDERS FOR DIABETIC KETOACIDOSIS (DKA) INPATIENT AND OUTPATIENT

I	NPATIENT AN	ND OUTPATIENT				
	// O MM YYYY	TIME:HOURS				
WEIGHT_	kilograms	HEIGHT centimetres	☐ ALLERGY CAUTION sheet reviewed			
Pharmacy Use Only	WRITE FIRMLY WITH A BALLPOINT PEN WITH BLUE OR BLACK INK					
Use Only	On Admission STA  vital signs ar weigh patien strictly monit nothing by m pulse oximet insert large-k capillary blood urine for keto venous blood calcium, glud urea, creatin other labs:  Fluid Resuscitatio  1st: sodium of 2nd: sodium of 2nd: sodium of Fluid Repair (after Bag A: sodium determined f  Fluid Repair and I reduce Bag Bag B: dextra  8, to keep glu 0.9% at mL/kg/hour = mL of prepar  Ongoing Monitoria capillary glud venous blood calcium, glud if patient dev notify MD ST mannitol 20%	AT:  Ind neurovital signs on admission and the tor input and output and output and cardiac monitor pore intravenous cannula and glucose by fingerpoke ones and gas; whole blood sodium, potassium, and cose, beta-hydroxybutyrate ine, phosphorus, complete blood-cell complete blood-cell complete ones and gas; whole blood sodium, potassium, and cose, beta-hydroxybutyrate ine, phosphorus, complete blood-cell complete ones are initial 30–60 minutes):  In Bolus(es) (initial 30–60 minutes):  In Il V over 30 ml IV over 30 ml	chloride, bicarbonate, anion gap, ionized bunt/differential, HbA1C  minutes (10 mL/kg) minutes (10 mL/kg) minutes (10 mL/kg) minutes (10 mL/kg)  meq/L potassium chloride at arte determined from DKA protocol, line of DKA protocol, line 7, where 1 ding sites by priming and flushing with 50 and discard.  Sted 30–60 minutes) chloride, bicarbonate, anion gap, ionized reatinine, and phosphorus every  vital signs or Glasgow Coma Scale Score:  IV fluids to 5 mL/hour pending MD review 0.5–1 g/kg, 2.5–5 mL/kg)	RN/UC		



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http://endodiab.bcchildrens.ca

#### BCCH DKA RECIPES FOR MAKING SOLUTIONS

Item #		Dextrose solution to prepare:	1 Dextrose solution and size of IV bag to use:	2 Withdraw & discard from bag:	3 Add to bag:		
1	commercially available	D5W-NaCl 0.9% with 40 mmol KCl /L					
2		<b>D10W</b> -NaCl 0.9% with 40 mmol KCl /L	1000 mL D5W-NaCl 0.9% with 40 mmol KCl /L	100 mL	100 mL <b>D50W</b>		
3		D12.5W-NaCl 0.9% with 40 mmol KCl /L	1000 mL D5W-NaCl 0.9% with 40 mmol KCl /L	100 mL	150 mL <b>D50W</b>		
4	commercially available	D5W-NaCl 0.45% with 40 mmol KCl /L					
5		D10W-NaCl 0.45% with 40 mmol KCl /L	1000 mL D5W-NaCl 0.45% with 40 mmol KCl /L	100 mL	100 mL <b>D50W</b>		
6		D12.5W-NaCl 0.45% with 40 mmol KCl /L	1000 mL D5W-NaCl 0.45% with 40 mmol KCl /L	100 mL	150 mL <b>D50W</b>		
Note: this	Note: this results in approximate concentrations and is to be used only when Pharmacy mixing is not available						
Prepared	Prepared by C&W Pharmacy Department; contact 604-875-2059 for questions						



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#### BCCH DKA GLUCOSE, INSULIN AND FLUID MANAGEMENT

#### 2019 REVISIONS TO BCCH DKA PROTOCOL

The 2019 revisions to the BCCH DKA Protocol are based on the results of recent research findings on rehydration protocols. These revisions brings the BCCH DKA Protocol into alignment with the *Clinical Practice Consensus Guidelines 2018* from the International Society for Pediatric and Adolescent Diabetes (ISPAD) and with the 2018 DKA resources from TREKK Canada (references below).

#### INITIAL FLUID REPLACEMENT

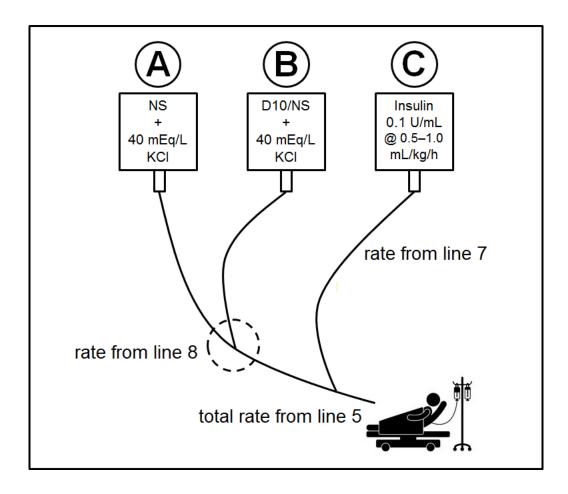
Results from the PECARN DKA FLUID Study (reference below) have demonstrated that fluid replacement can safely be achieved using more-aggressive regimens than have been in place over the past two decades. It is now recommended that all patients in moderate-to-severe DKA receive a 10-mL/kg bolus of 0.9% sodium chloride (normal saline, NS) over 30 minutes. Those patients with persistent tachycardia, prolonged capillary refill (>2 sec), and cool extremities should receive a second 10-mL/kg fluid push as well. Once the fluid push(es) have been delivered, and assuming the patient has adequate urine output and a normal serum potassium, fluid replacement is continued using NS + 40 mEq/L KCl (Bag A, see next section), until the patient has been receiving fluids for 2 hours; at that point, intravenous insulin is started. Fluid replacement rates are now calculated for a 36-hour period of rehydration, compared to the 48-h period used in the past.

#### THE "TWO-BAG SYSTEM"

The "two-bag system" (reference below) consists of two IV bags ( $\mathbf{A}$  and  $\mathbf{B}$ ) with equal electrolyte concentration, one containing no dextrose, the other 10-12.5% dextrose. They are administered simultaneously. The total rate is determined by the child's degree of dehydration, according to the BCCH DKA Medical Protocol (line 5). The insulin infusion ( $\mathbf{Bag}\ \mathbf{C}$ ) will eventually be  $\mathbf{Y}$ 'd into these bags (see below).

In the "two-bag system", Bag A is generally NS + 40 mEq/L KCl, and Bag B is usually D10/NS + 40 mEq/L KCl (or D12.5/NS + 40 mEq/L KCl, if your institution can make this). The BCCH Pharmacy has prepared a "recipe book" for preparing these solutions from

commercially available IV solutions, which is available from the Parenteral Drug Manual on the BCCH ePOPS website and on BCCH Endocrinology's website (references below).



MANAGING THE BLOOD GLUCOSE (BG) LEVEL

The goal is to keep the BG levels in the 8-12 mmol/L range, both to minimize glycosuria and to allow for a buffer against hypoglycemia. This is most easily achieved by alternately adjusting the rates of the non-dextrose-containing Bag A and the dextrose-containing Bag B, while keeping the insulin infusion rate constant (see example below).

At the onset, it is recommended having both bags prepared and hung, starting Bag A at nearly the full rate (from line 8 of the BCCH DKA Medical Protocol), and starting Bag B at a "to-keep-open" rate (2-5 mL/h). The insulin infusion rate of 0.05-0.1 U/kg/h (0.5-1.0 mL/kg/h) should not be adjusted until the pH is close to normal (see below).

Example: IV rate from protocol line 8 = 100 mL/h (this does not include insulin infusion rate) rate Bag A + rate Bag B = 100 mL/h

rate Bag A no dextrose (mL/h)	rate Bag B D10 (mL/h)	final dextrose concentration (%)
100	0	D0
80	20	D2
60	40	D4
40	60	D6
20	80	D8
0	100	D10

The BG level will fall quite rapidly in the first hour or two with the initial fluid management, even before insulin is started, secondary to improved renal clearance and hemodilution. Thereafter, one should aim for a fall in BG of  $\sim$ 3-5 mmol/L/h.

Assuming that the BG is being monitored every 30-60 minutes, once it starts to approach ~20 mmol/L—sooner if the initial BG drop is >5 mmol/L/h—the rate of Bag A is decreased, and the rate of Bag B is increased by an equivalent amount. A general rule is to make changes of approximately 10-20% of the total every hour. This will depend on the rate of fall of the BG level and the patient's response to these changes.

If the patient's BG level is lower than desired, despite maximal dextrose infusion from Bag B, you may (in order of safety):

- 1. cut the insulin infusion rate by ~25%, provided the acidosis is correcting
- 2. give the patient a small amount (1-2 mL/kg) of juice or 2-4 dextrose tablets (being mindful of the overall fluid balance)
- 3. change the insulin bag to D10/NS
- 4. in institutions with intensive-care capabilities, consider placing a central line and using a higher concentration of dextrose (e.g. D20) in Bag B.

#### THE INSULIN INFUSION

The optimal initial insulin infusion rate is not known, but an increasing number of experts are suggesting a starting rate of 0.05 U/kg/h, i.e. 50% of the rate of previous protocols. ISPAD 2018 (reference below) supports the use of either starting rate, until more conclusive information is available. We would suggest that this lower rate be considered especially when (1) patients have already had a significant drop in their BG prior to starting insulin; (2) when the patient's acidosis is less severe; (3) or when it is expected that the patient will be quite insulin-sensitive (some young children with DKA, patients with hyperglycemic hyperosmolar state, and some older children with established diabetes and insulin pump-site failure or acute insulin omission).

The half-life of IV insulin is quite short (minutes), so the insulin infusion should never be discontinued, until the patient has been established on subcutaneous insulin. If the patient's BG level is difficult to maintain >8-10 mmol/L despite the measures suggested above, one can cut the insulin infusion rate by  $\sim25\%$ , provided that the metabolic acidosis is resolving. It is unusual for a child in DKA to need <0.025 U/kg/h.

#### POTASSIUM

Nearly all children in DKA will require large amounts of potassium for repletion, and 40 mEq/L KCl in the IV will generally suffice. Some children will require extra oral or nasogastric potassium chloride (0.5-1.0 mEq/kg) to keep their serum potassium level >3.5 mmol/L. Rarely, children will require less potassium, in which case one could use 20 mEq/L.

#### SWITCHING TO HALF-NORMAL SALINE

The goal of treating DKA is to slowly allow the BG and hyperosmolality to normalize, which initially requires the use of isotonic fluids, i.e. normal (0.9%) saline. After about 4-6 hours, once the corrected  $Na^+$  is  $\ge 145$  mmol/L, the patient may require some free water in the form of hypotonic fluids to continue to have a drop in serum osmolality. At this point, Bags A and/or B can be switched to their half-normal (0.45%) saline equivalents.

#### HELP IN REAL TIME

If you have questions or problems related to the management of DKA or diabetes (for patients in BC and the Yukon), please feel free to contact the BC Children's Hospital Pediatric Endocrinologist on call at 604-875-2161.

#### ONLINE LINKS

The following resources are all available on our BCCH DKA Protocol webpage:

- BCCH DKA Protocol Toolkit
- BCCH DKA Medical Protocol (PLAIN PDF FORMAT)
- BCCH DKA Medical Protocol (FILLABLE PDF FORMAT)
- BCCH DKA Nursing Protocol
- BCCH DKA Flowsheet
- BCCH DKA Sample Prescriber Order Sheet
- BCCH DKA Glucose, Insulin and Fluid Management
- BCCH DKA Recipes for Making Solutions
- Blood Glucose and Insulin Record for Conventional Insulin Regimens
- Blood Glucose and Insulin Record for MDI

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Translating Emergency Knowledge for Kids (TREKK Canada): trekk.ca.

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BC Children's Hospital SHOP (Shared Health Organizations Portal): shop.healthcarebc.ca/phsa/bc-cnw-hospitals.