



Current Controversies in the Management of Hyperkalemia

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- The program chair and presenters for this continuing education activity have reported no relevant financial relationships.

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Blog post summarizing key points from today's presentation

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Debates in the Management of Hyperkalemia: A Focus on Calcium and Insulin

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A photograph of a patient's arm in a hospital setting. The arm is resting on a white surface. A red arterial cannula is inserted into the arm, secured with white tape. Several red monitoring wires are wrapped around the arm. In the background, a white sink and a yellow table are visible.

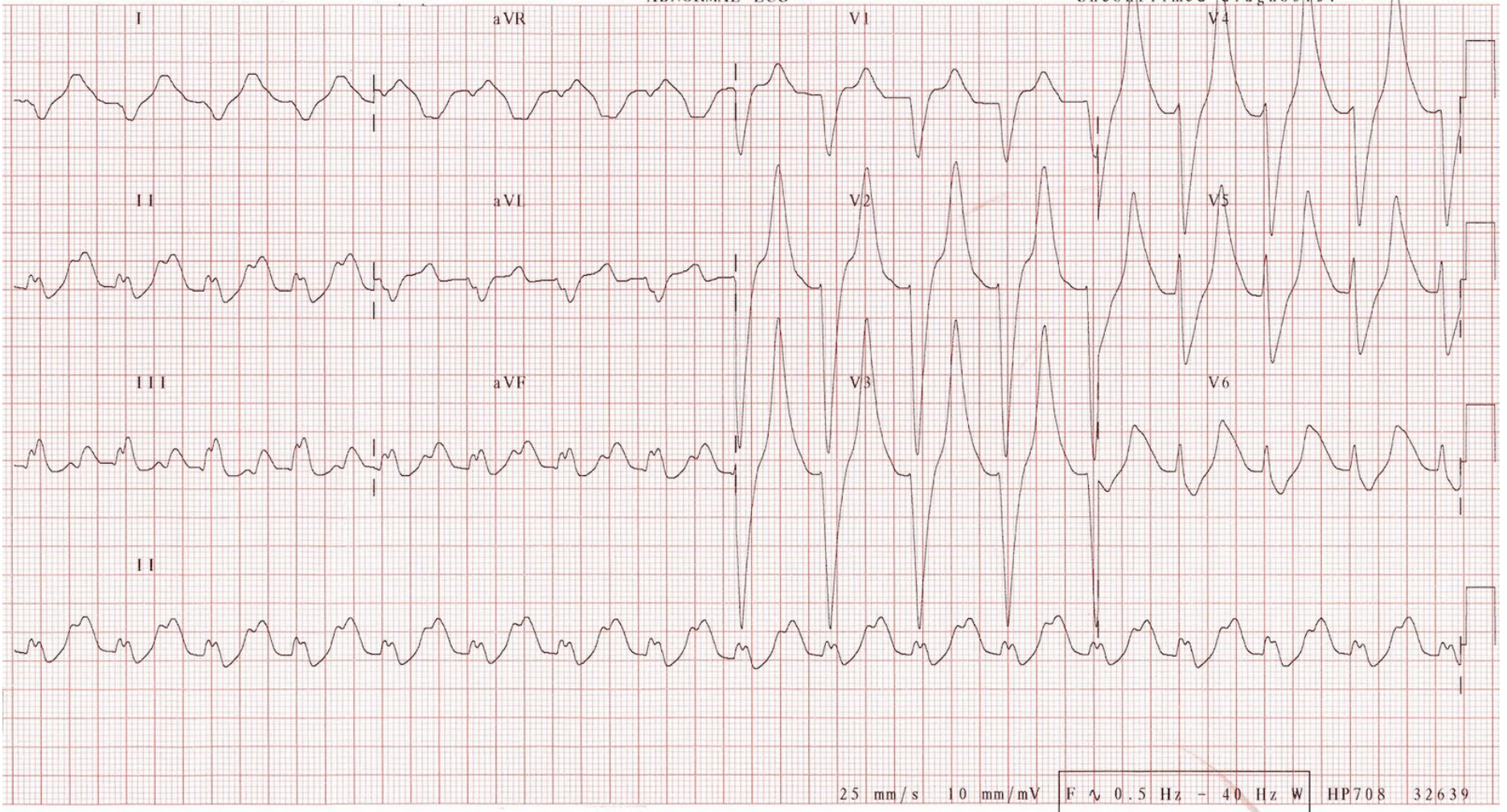
AMS

84/49 mm Hg

51 bpm

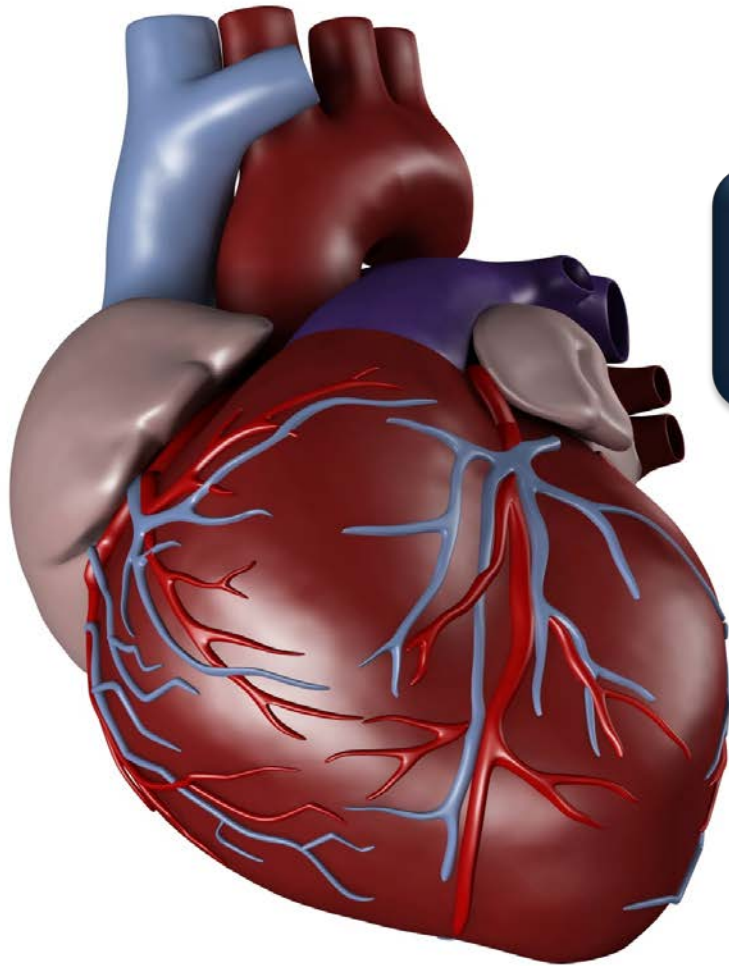
- ABNORMAL ECG -

Unconfirmed diagnosis.



Ca





↓ depolarization

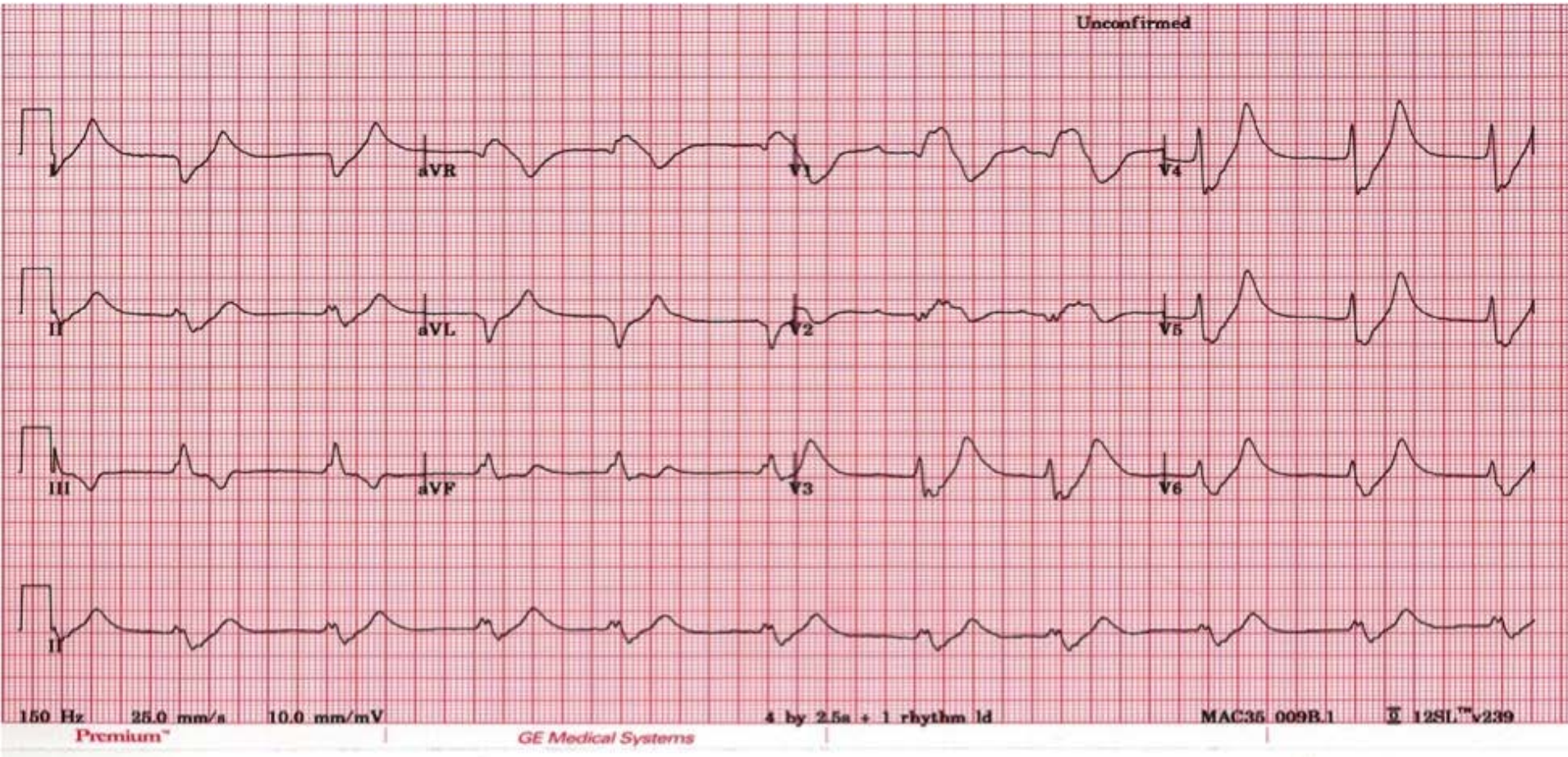
At what potassium concentration should calcium be administered?

- A 9.0 mmol/L
- B 7.1 mmol/L
- C 5.3 mmol/L
- D It depends

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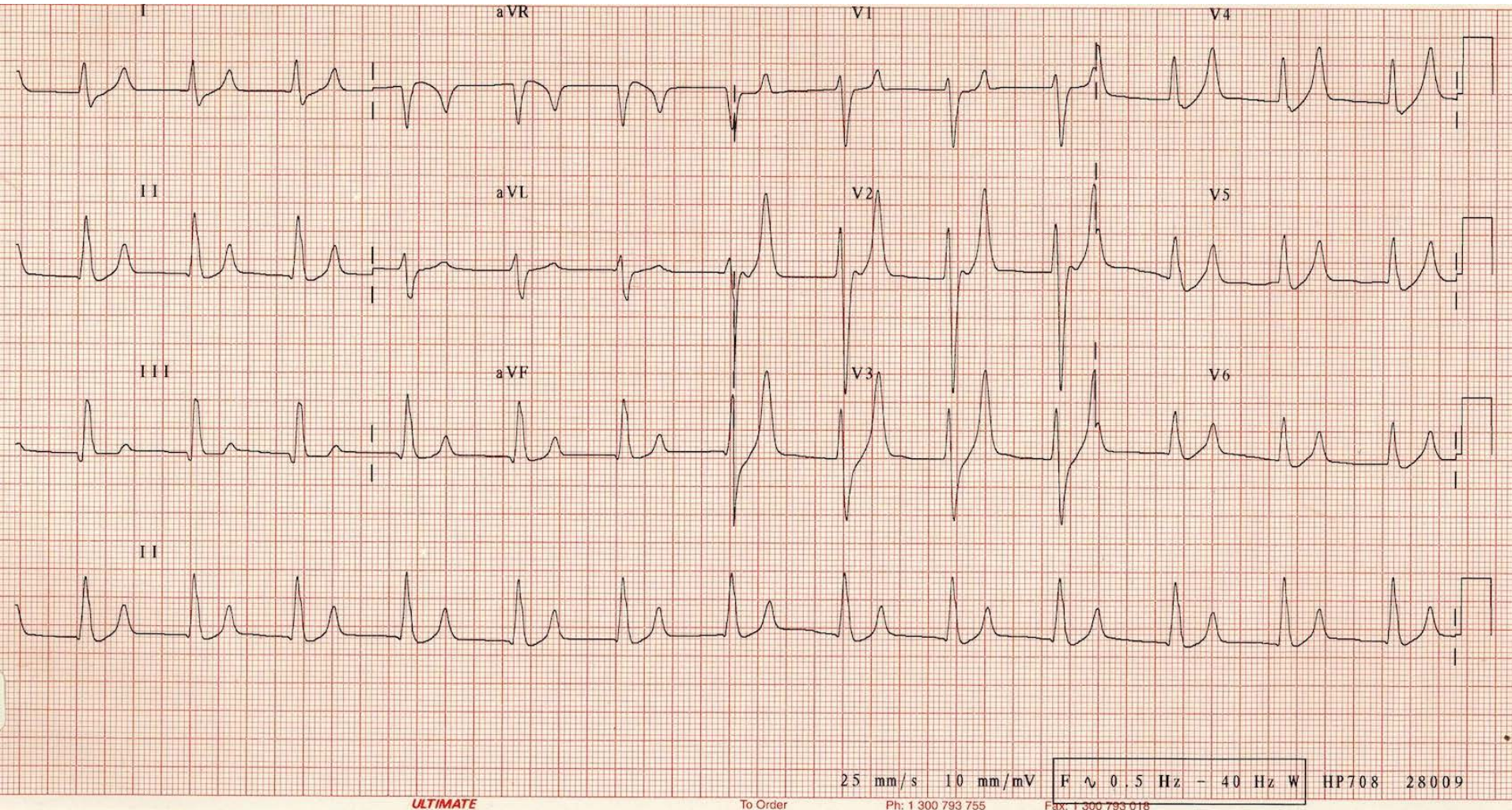
K⁺ 9.0 mmol/L



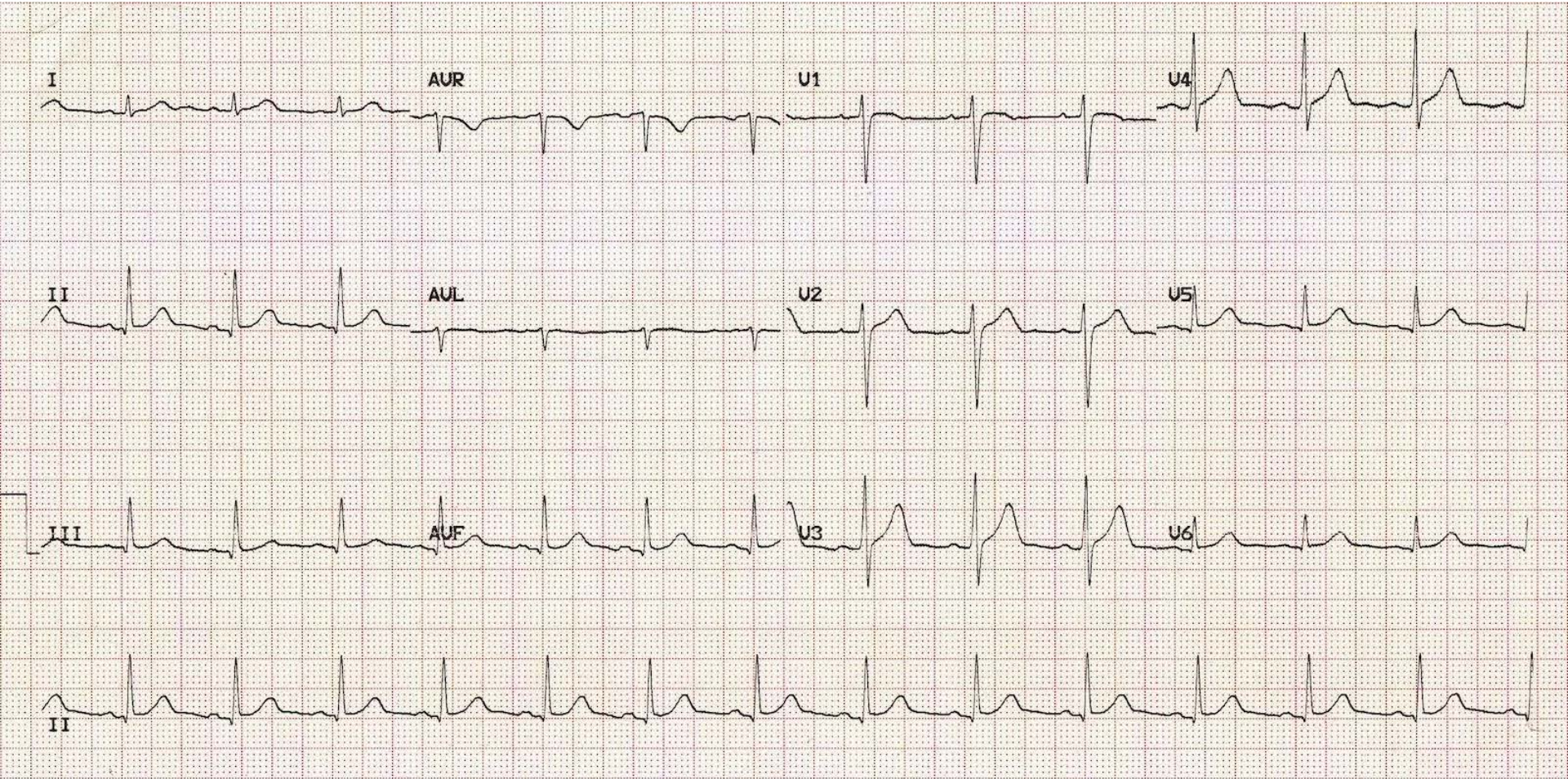
ECG courtesy of Life in the Fast Lane



K⁺ 7.1 mmol/L



K⁺ 5.3 mmol/L



ECG courtesy of Life in the Fast Lane



≥ 2 gm



≥ 1 gm

Which calcium salt works faster?

- A Calcium chloride
- B Calcium gluconate



Onset: 3 min

Duration: 30 min



Anesthesiology
73:62-65, 1990

*Ionization and Hemodynamic Effects of Calcium Chloride and
Calcium Gluconate in the Absence of Hepatic Function*

Thomas J. Martin, M.D.,* Yoogoo Kang, M.D.,† Kerri M. Robertson, M.D., F.R.C.P.(C.),*
Mohamed A. Virji, M.D., Ph.D.,† Jose M. Marquez, M.D.†

Equal rise in ionized Ca^{2+}

Anesthesiology
66:465-470, 1987

Calcium Chloride Versus Calcium Gluconate: Comparison of Ionization and Cardiovascular Effects in Children and Dogs

Charles J. Cote', M.D.,* Lambertus J. Drop, M.D.,† Alfred L. Daniels, M.S.,‡ David C. Hoaglin, Ph.D.§

Equal rise in ionized Ca^{2+}



Equal rise in ionized Ca^{2+}

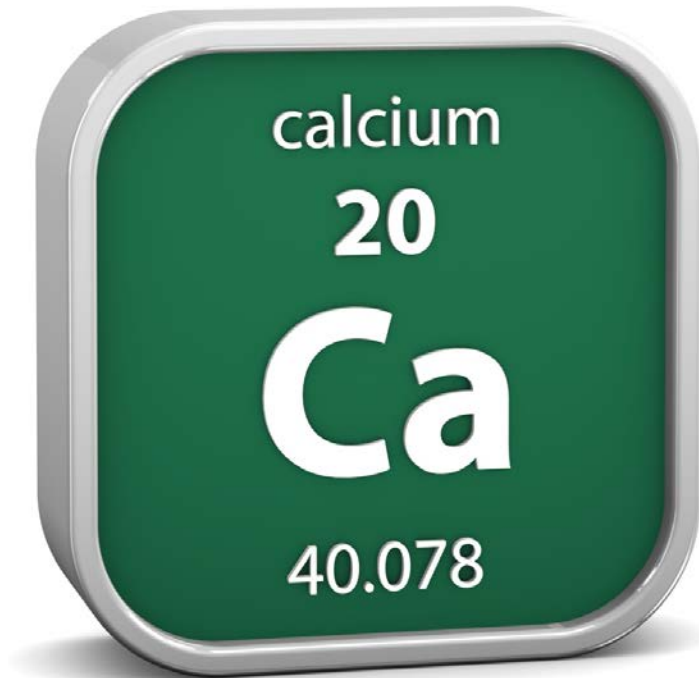
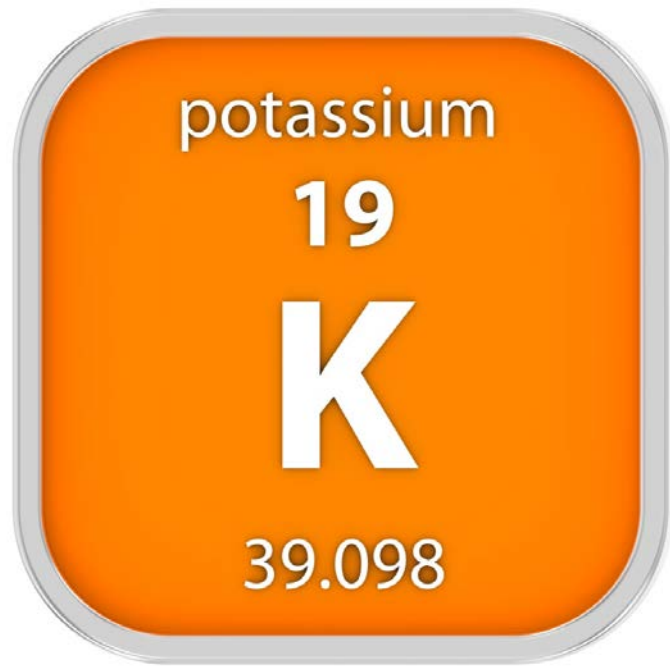
CaCl₂ NOT faster



Which calcium salt works faster?

- A Calcium chloride
- B Calcium gluconate**









5 cases

**Flawed
animal
models**

The effects of IV calcium in patients with digoxin toxicity.

Levine M, et al. *J Emerg Med* 2011;40(1):41-6.

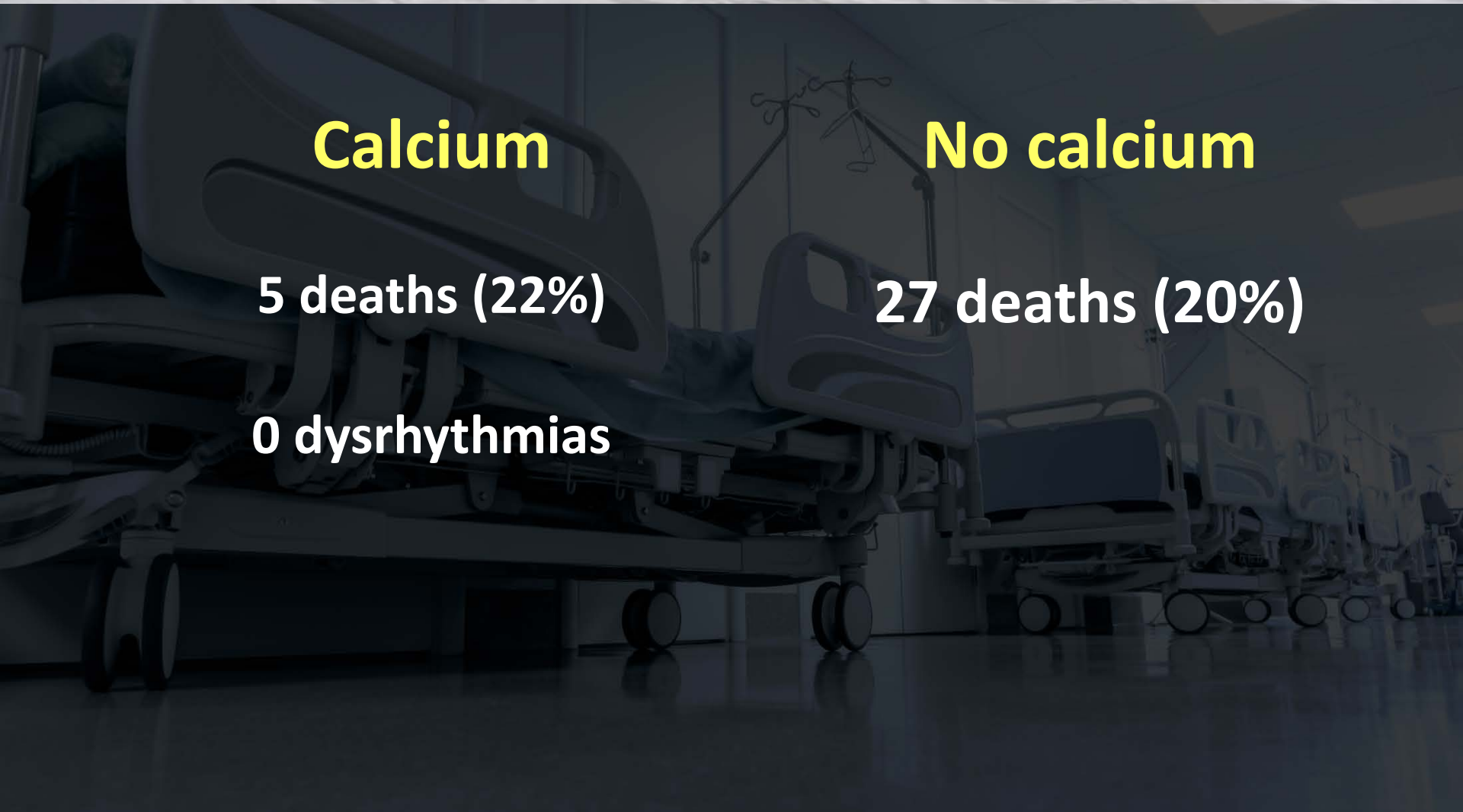
Calcium

5 deaths (22%)

0 dysrhythmias

No calcium

27 deaths (20%)







NDC 59060-1833-2

Novolin® **R**

Human Insulin
(recombinant
origin) USP
Units/mL 10 mL

(01) 103 5906 0183 32 9

Novo Nordisk®
Important: see insert
Keep in a cold place

NDC 59060-1834-2

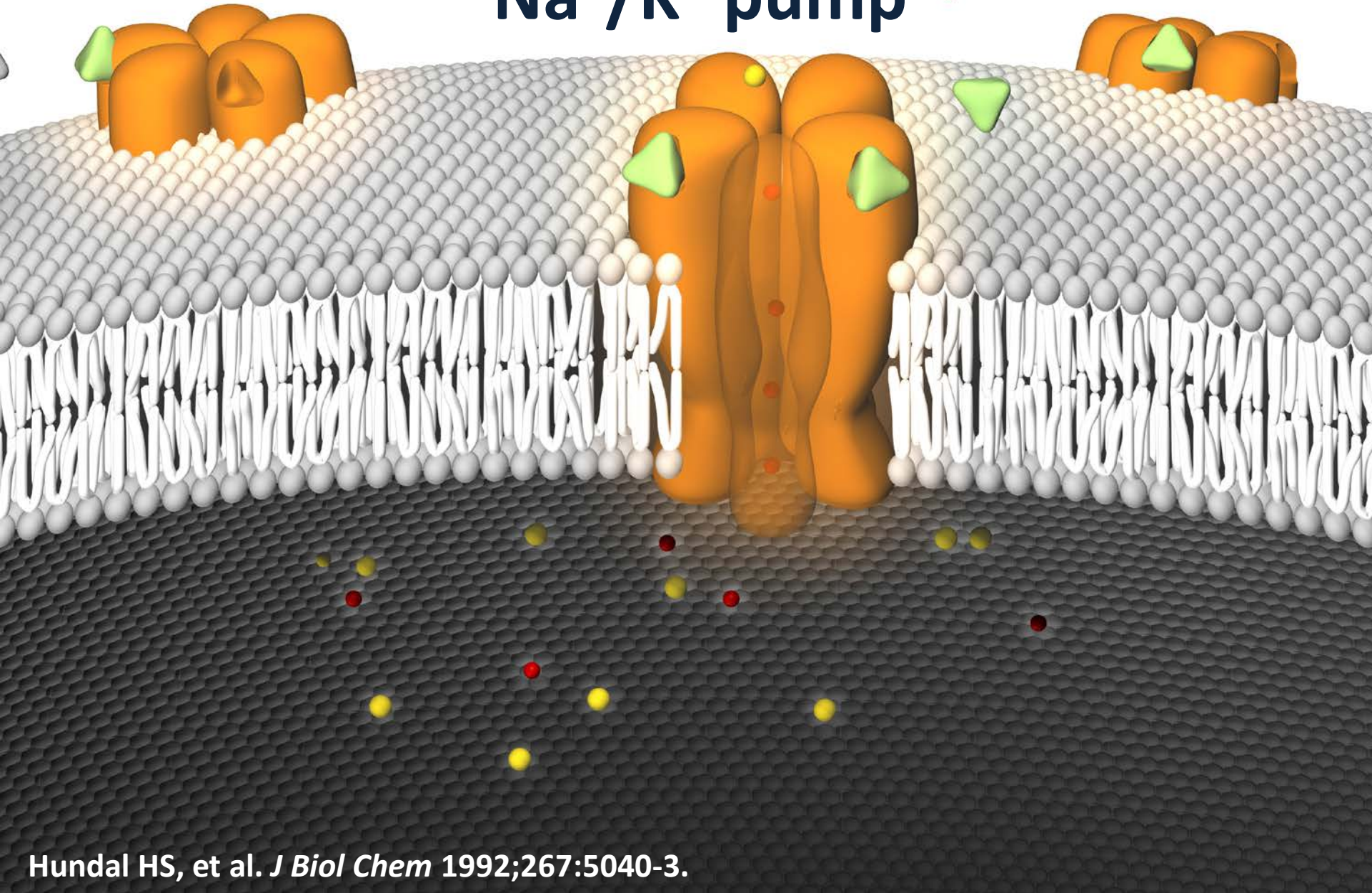
Novolin® **N**

Human Insulin
Suspension
(recombinant DNA)
Units/mL 10 mL

(01) 103 5906 0183 42 8

Novo Nordisk®
Important: see insert
To mix, shake carefully
Keep in a cold place

Na⁺/K⁺ pump

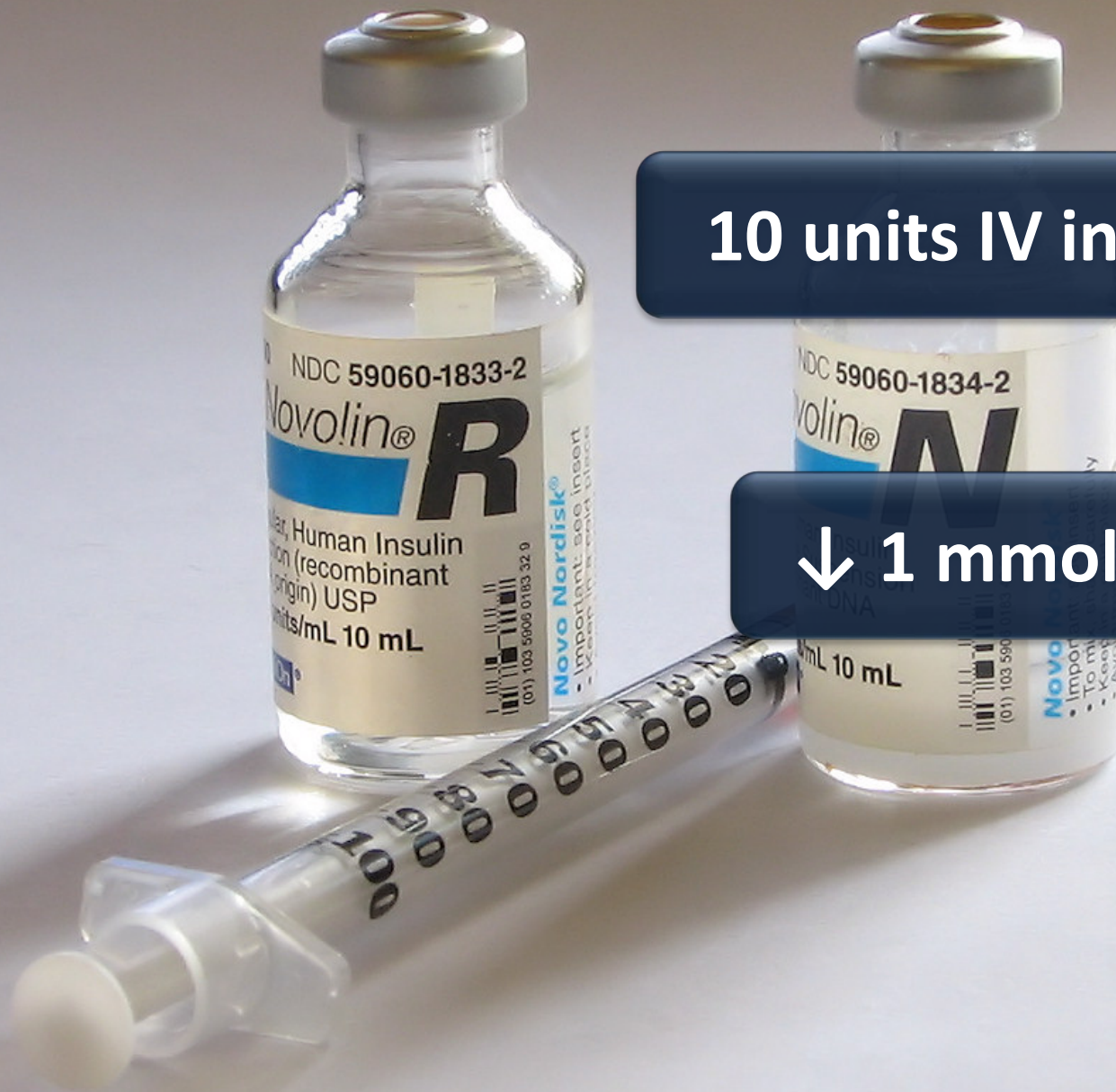


What is the most common insulin dose used for hyperkalemia?

- A 5 units IV
- B 10 units IV
- C 20 units IV
- D 20 units/hr IV

What is the most common insulin dose used for hyperkalemia?

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- B 10 units IV**
- C 20 units IV
- D 20 units/hr IV



10 units IV insulin

↓ 1 mmol/L



~ 10%

Schafers S, et al. *J Hosp Med* 2012;7(3):239-42.
Allon M, et al. *Kidney Int* 1990;38(5):869-72.

Onset

5-10

Peak

25-30

Duration

120-180



30-60 min



No DM

No DM meds

**Lower glc
(104 vs 162)**

Renal

**Risk
Factor**

Dickerson RN, et al. *Nutrition* 2011;27:766-72.

Apel J, et al. *Clin Kidney J* 2014;0:1-3.

Initial [glucose]	Initial Dextrose Dose	Supplemental Dextrose **	Glucose Monitoring
> 200 mg/dL (> 11.1 mmol/L)	None	50 mL (25 gm) of D50 if blood glucose < 70 mg/dL	Hourly up to 3 hours
100-200 mg/dL (5.6-11.1 mmol/L)	50 mL (25 gm) of D50	50 mL (25 gm) of D50 if blood glucose < 70 mg/dL	Hourly up to 3 hours
< 100 mg/dL (< 5.6 mmol/L)	100 mL (50 gm) of D50* OR 50 mL (25 gm) of D50 + D10 infusion 250 mL/hr for first hour	50 mL (25 gm) of D50 if blood glucose < 70 mg/dL	q 30 minutes for first hour, then hourly up to 3 hours

D50 = dextrose 50%; D10 = dextrose 10%

Summary

- 1 ECG changes? Give calcium
- 2 Calcium gluconate = CaCl_2
- 3 Dig toxicity? Give antidote (+ calcium)
- 4 Anticipate hypoglycemia



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Debates in the Management of Hyperkalemia: Sodium “Redux” and Novel Therapies

Nadia I. Awad, PharmD, BCPS

Emergency Medicine Pharmacist

Robert Wood Johnson University Hospital



@Nadia_EMPHarmD

Sodium bicarbonate is frequently administered to hyperkalemic patients in my institution.

A Yes

B No

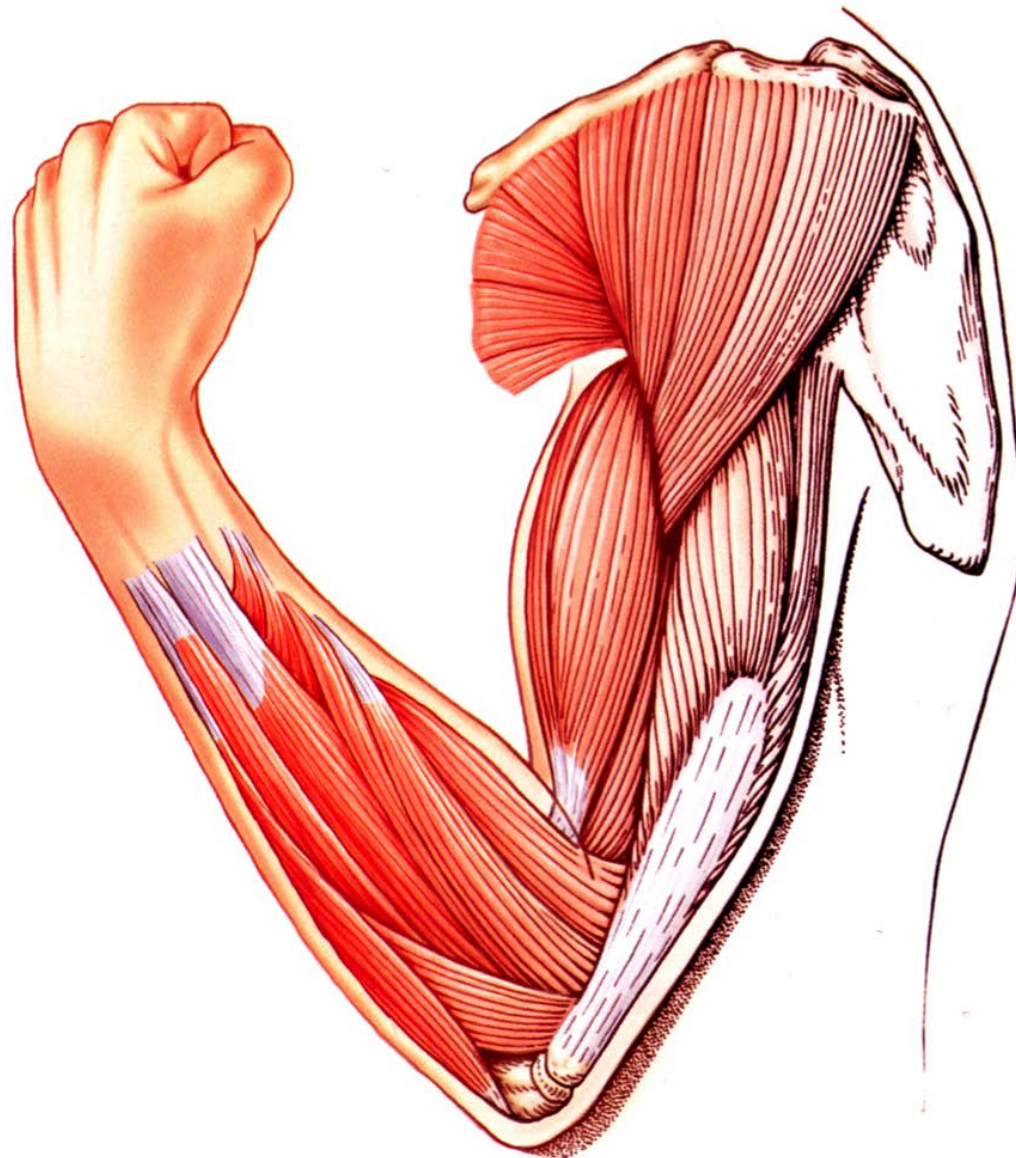
Preferred Therapy of Hyperkalemia in Renal Insufficiency: Survey of Nephrology Training-Program Directors

N Engl J Med 1989; 320:60-61.



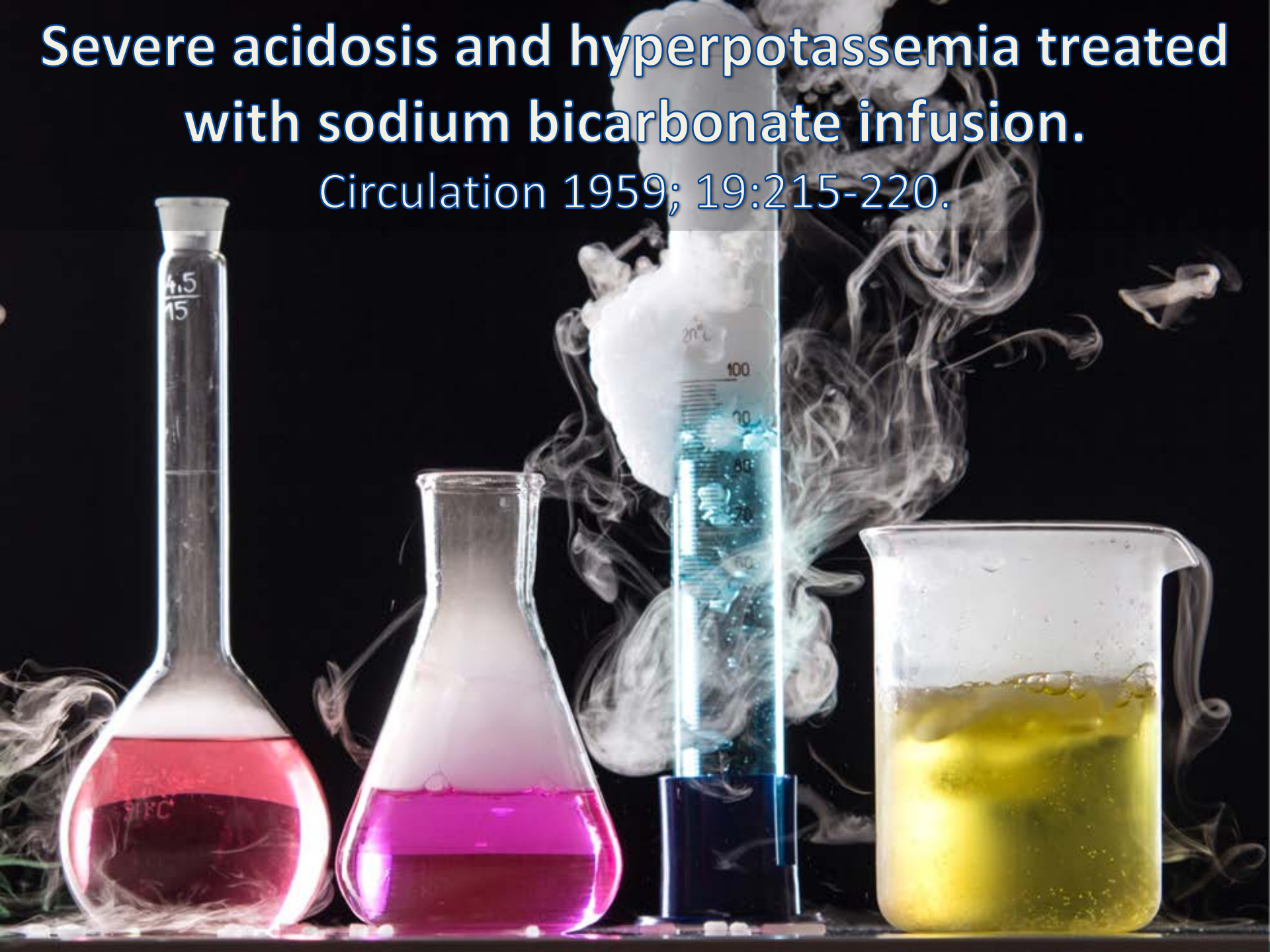
Sodium bicarbonate works in hyperkalemia because it:

- A Enhances K^+ excretion
- B Shifts K^+ to the intracellular space
- C Decreases H^+ in the extracellular fluid
- D Works some other way that I learned about but cannot recall right now



Severe acidosis and hyperpotassemia treated with sodium bicarbonate infusion.

Circulation 1959; 19:215-220.





Am J Med 1988; 85:507-512.
Nephron 1996; 72:476-482.
Am J Kidney Dis 1996; 28:508-514.
Korean Med Sci 1997; 12:111-116.



Miner Electrolyte Metab 1991; 17:297-302.
Kidney Int 1992; 41:369-374.

#WBW



J Am Med Assoc 1954; 155:877-883.
Am J Kidney Dis 1996; 28:508-513.
East Afr Med J 1997; 74:503-509.

“Given this uncertainty we still use sodium bicarbonate to treat acute hyperkalemia in patient with a significant degree of acidosis, but not as the only emergency therapy.”

Nephrol Dial Transplant 2003; 18:2215-2218.



“We suggest that intravenous sodium bicarbonate infusion is not used routinely for the acute treatment of hyperkalemia.”

Hyperkalemia Guidelines, UK Renal Association 2014.

Management of severe hyperkalemia

Lawrence S. Weisberg, MD

Crit Care Med 2008; 36:3246-3251.

Citations: 130 (as of October 2016)

Am Heart J 1962; 64:483-488.

Am J Med 1953; 14:504.

Acad Emerg Med 1997; 4:93-99.

Acad Emerg Med 2000; 7:965-973.

Code Name: Nav1.5

Depolarization

Cellular
perfusion

Afterload

Dilution



Am J Physiol 1975; 229:935-940.
Acad Emerg Med 2000; 7:965-973.
Kidney Int 2016; 90:450-451.

The dose of sodium polystyrene sulfonate routinely given for hyperkalemia within my institution is:

A 0 g

B 15 g

C 30 g

D 60 g

The use of polystyrene sulfonate in the inpatient management of hyperkalemia.

J Hosp Med 2011; 6:136-140.

10'

0.82 to 1.4 mEq/L

1958

1962

Rx Only

NDC 0574-2003-16

**Sodium
Polystyrene
Sulfonate
Suspension,
USP**

15 g/60 mL

Does not contain Sorbitol

Dispense in tight container.

SHAKE WELL BEFORE USING

Protect from freezing and from
excessive heat.

FOR ORAL OR RECTAL USE

473 mL (one pint)



Lancet 1953; 265:791-795.

Gastroenterology 1994; 107:548-571.

N Engl J Med 1961; 264:111-115. (n = 1)
N Engl J Med 1961; 264:115-119. (n = 32)






100 mg Na per g SPS

Aldosterone and ESRD

2009: Cathartic controversy: 33% / 70%?

Time / Replication

Am J Kidney Dis 1988; 8:105-110.
Gastroenterology 1995; 108:752-760.
J Am Soc Nephrol 1998; 9:1924-1930.
Am J Med 2013; 126:264.e9-264.e24.
J Am Soc Nephrol 2010; 21:733-735.

A glowing blue hammer is shown striking a metal rod, creating a bright yellow and orange spark. The hammer and rod are illuminated with a blue glow, and the background is dark. The text is overlaid on the image.

“In summary, we do not use resins for treatment of acute hyperkalemia. In the setting of chronic hyperkalemia, it seems that the addition of resins to cathartics adds little to the induction of diarrhea alone.”

Nephrol Dial Transplant 2003; 18:2215-2218.

I believe the novel therapies for hyperkalemia (patiromer and sodium zirconium cyclosilicate) may have a role in treating hyperkalemia.

A Yes

B No

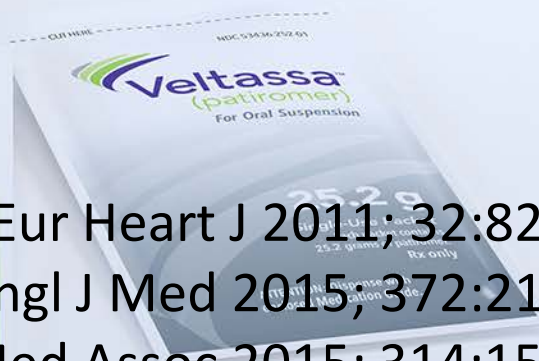
“Me too”

CKD and CV

Long-term

BBW

Mg



Eur Heart J 2011; 32:820-828.

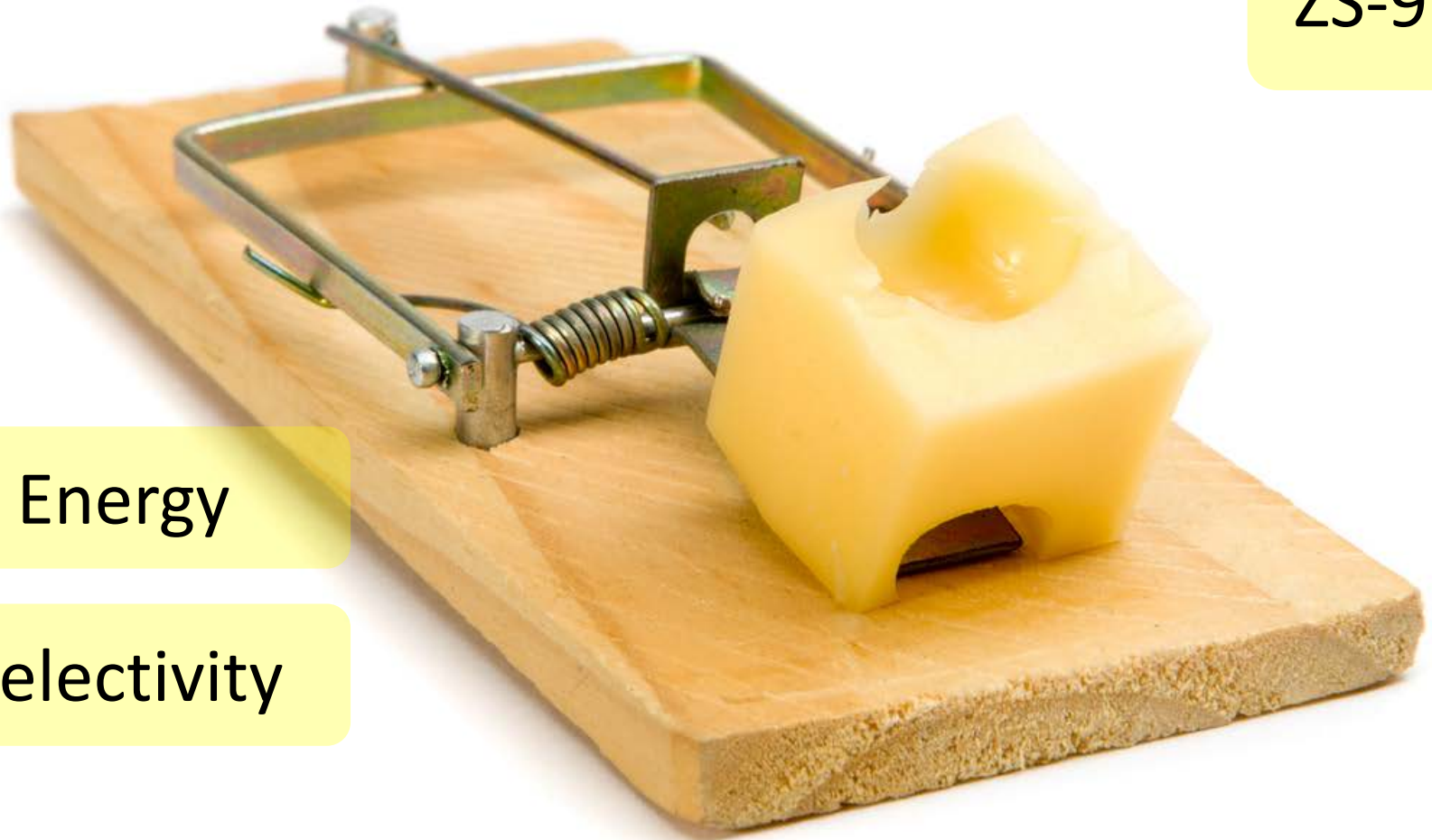
N Engl J Med 2015; 372:211-221.

J Am Med Assoc 2015; 314:151-161.

ZS-9

Energy

Selectivity



PLoS One 2014; 9:e114686.
Kidney Int 2015; 88:404-411.
N Engl J Med 2015; 372:222-231.

Sodium Zirconium Cyclosilicate for Urgent Therapy of Severe Hyperkalemia.

N Engl J Med 2015; 372:1577-1578.

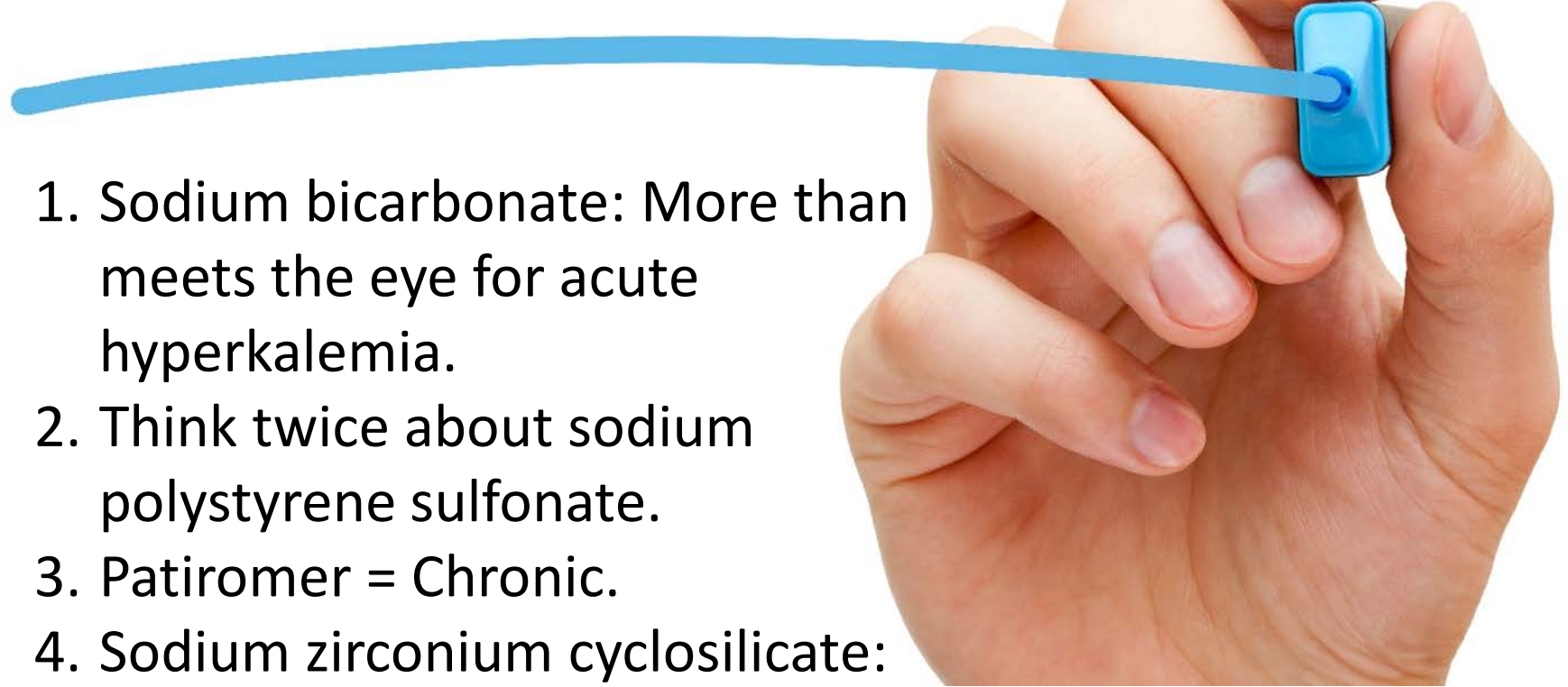
1'	0.4 mmol/L
2'	0.6 mmol/L
4'	0.7 mmol/L

Serum K 6.1 to 7.2 mmol/L + 10g ZS-9



Kidney Int 2015; 88:404-411.
N Engl J Med 2015; 372:222-231.
Hypertension 2015; 66:731-738.
Pharmacotherapy 2016; 36:922-923.

SUMMARY



1. Sodium bicarbonate: More than meets the eye for acute hyperkalemia.
2. Think twice about sodium polystyrene sulfonate.
3. Patiromer = Chronic.
4. Sodium zirconium cyclosilicate: Don't jump on the bandwagon.



**Debates in the Management of
Hyperkalemia:
Sodium “Redux” and Novel Therapies**

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