Paisley Ties, Robert Plant, and Sodium Bicarbonate

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The Plan

Review evidence regarding sodium bicarbonate

Experience in an animal model

What should we actually be doing?





Ivory Tower

In the Streets



Parachute use to prevent death and major trauma related to gravitational challenge: systematic review of randomised controlled trials

Gordon C S Smith, Jill P Pell



Parachutes reduce the risk of injury after gravitational challenge, but their effectiveness has not been proved with randomised controlled trials

What is already known about this topic

Parachutes are widely used to prevent death and major injury after gravitational challenge

Parachute use is associated with adverse effects due to failure of the intervention and iatrogenic injury

Studies of free fall do not show 100% mortality

What this study adds

No randomised controlled trials of parachute use have been undertaken

The basis for parachute use is purely observational, and its apparent efficacy could potentially be explained by a "healthy cohort" effect

Individuals who insist that all interventions need to be validated by a randomised controlled trial need to come down to earth with a bump

What We Know

Large clinical trial with post-hoc analysis

Animal models with fast sodium channels

→ Jim Menegazzi's cocktail



Improved resuscitation outcome in emergency medical systems with increased usage of sodium bicarbonate during cardiopulmonary resuscitation

G. Bar-Joseph^{1,4}, N. S. Abramson^{1,2}, S. F. Kelsey³, T. Mashiach⁵, M. T. Craig¹ and P. Safar¹ for the Brain Resuscitation Clinical Trial III (BRCT III) Study Group

¹Safar Center for Resuscitation Research, and Departments of ²Emergency Medicine and ³Epidemiology, University of Pittsburgh, Pittsburgh, PA, and ⁴Pediatric Intensive Care Unit and ⁵Quality Assurance Department, Rambam Medical Center and the Bruce Rappaport Faculty of Medicine, Technion – Israel Institute of Technology, Haifa, Israel

What They Did

♣ This was the "high dose" epi trial

- Looked at 3 types of sodium bicarbonate users:
 - High users (5): Use bicarb in over 50% of cases and mean time to bicarb < 10 mins</p>
 - Intermediate users (3): One of two above
 - Low users (8): Less than 50% and mean time to bicarb >10 mins

	ROSC ROSC+/total, n (%)		
	Low SB users	High SB users	P*
VF or VT AS or PEA	151/578 (26.1) 115/448 (25.7)	91/281 (32.4) 141/413 (34.1)	0.056 0.007

Good neurological of CPC1 or 2/total, n (
Low SB users	High SB users	P*
16/577 (2.8) 6/448 (1.3)	23/279 (8.2) 14/443 (3.4)	0.0005 0.005

	ROSC rate			
Variable	OR	95% CI	P	
Time collapse to ACLS ¹	1.32	1.07-1.63	0.01	
Time from ACLS to Epi ²	1.17	0.94-1.46	NS	
Presenting rhythm ³	1.02	0.82-1.27	NS	
SB usage profile ⁴	1.36	1.08-1.70	0.007	
Epi treatment group ⁵	0.90	0.64-0.98	NS	
Epi treatment group ⁵ Sex ⁶	1.40	1.11-1.77	0.004	
Age ⁷	1.21	0.97-1.51	NS	

Odds of Neurologically Intact Survival

Time collapse to ACLS ¹	4.08	2.17-7.67	< 0.001
Time from ACLS to Epi ²	3.18	1.69-5.98	< 0.001
Presenting rhythm ³	2.75	1.52-4.98	0.001
SB usage profile4	2.18	1.23-3.86	< 0.008
Epi treatment group ⁵ Sex ⁶	1.11	0.65-1.89	NS
Sex ⁶	1.43	0.77-2.65	NS
Age ⁷	2.12	1.13-3.97	0.02

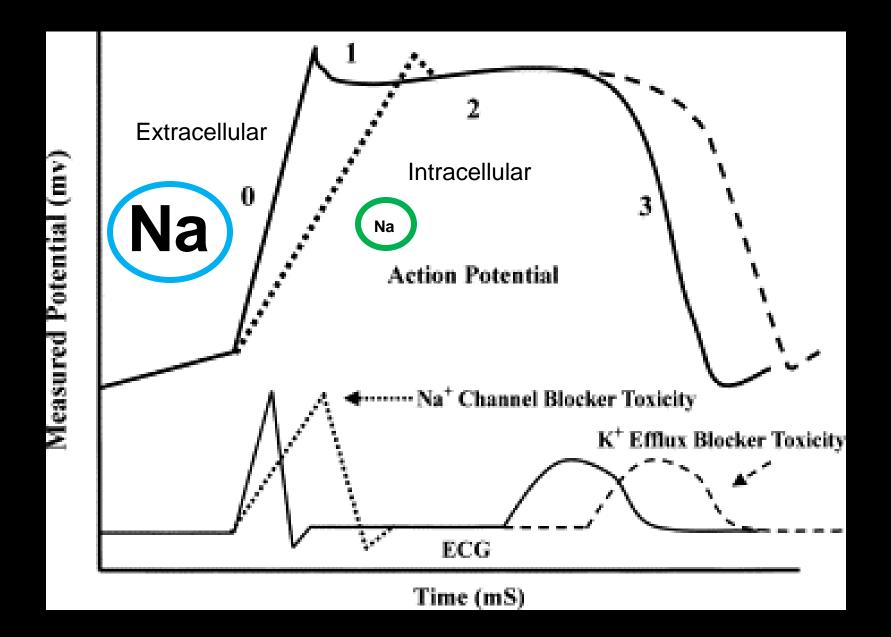
How Does It Work?

→ Acid/Base believers

Sodium load believers

Some combination





TCA overdose

QRS widens, presumably due to relative sodium channel blockade

Cocaine has a similar mechanism

Same effect (via different mechanism) for hypoxia-induced loss of extra vs. intra cellular sodium gradient

Animal Study

All animals we made toxic with TCA

- ♣ 4 treatment groups
 - → Control Just D5W
 - Hypertonic saline
 - → Sodium bicarbonate
 - Hyperventilation to achieve pH of 7.5 to 7.6

Table 3.

Survival at 60 minutes.

Group	Subjects Surviving		
Control	1/6	17%	
HTS	5/6	83%	
NaHCO ₃	2/6	33%	
Hyperventilation	1/6	17%	

Does The Same Hold True for Class I antidysrhythmic OD?

TABLE 1 -- Spontaneous VT Protocol: Flecainide Doses, Flecainide Plasma Concentrations, and Response to Treatment

Total flecainide dose at onset of VT (mg/kg)

Flecainide concentration 1 minute after third treatment (ng/mL)

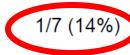
Termination of VT by treatment (n)

Time to recurrence of VT after treatment (min)

Control

 58 ± 14

7448 ± 2357



1.8 *

NaHCO₃

 57 ± 24

 5411 ± 2542



19.2 ± 24.4

[†] P < .01 v control.

^{*} SD omitted because only one animal was a responder.

Summary So Far

No randomized controlled trial

Best evidence in humans from one paper – seems to support bicarbonate use

Animal studies also support use in limited situations



What Should We Do?

Studies that placed bicarbonate in a lower priority were not of the highest quality

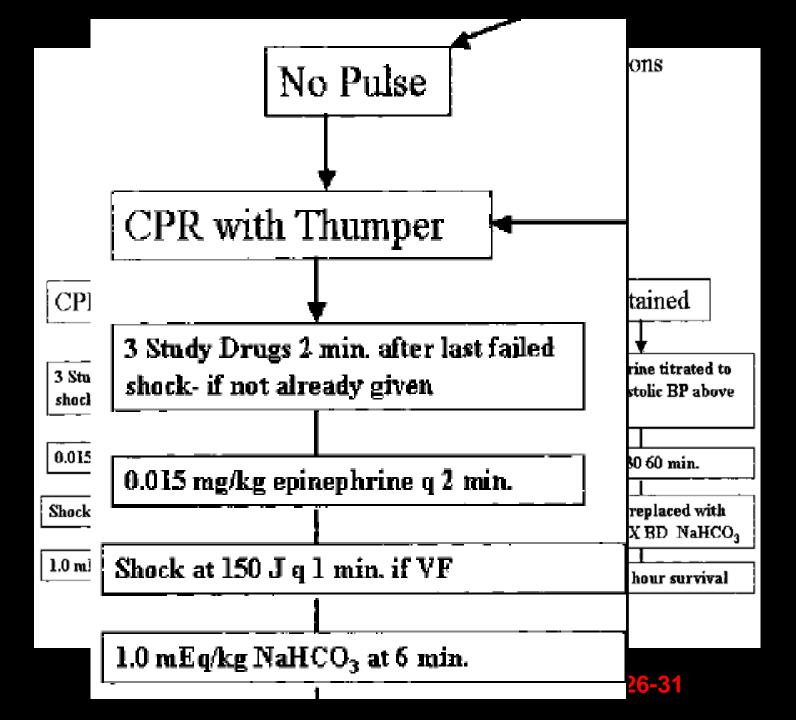
A modest proposal follows



Menegazzi's Protocol

- The protocol is not the point of the study
- The study needs animals to achieve ROSC
- "Three drugs" are vasopressin, epinephrine, and propanolol





When the Call Comes

- Give good CPR
- Give airways their due
- Give pressors
- If refractory VF/VT or PEA/Asystole, consider bicarb
- If cocaine, TCA, or other sodium channel blockade a concern, give bicarb early

