



# Squilibri Elettrolitici in PS: Differenze Età Correlate

Nuovi Indicatori di Performance in Medicina D'Urgenza

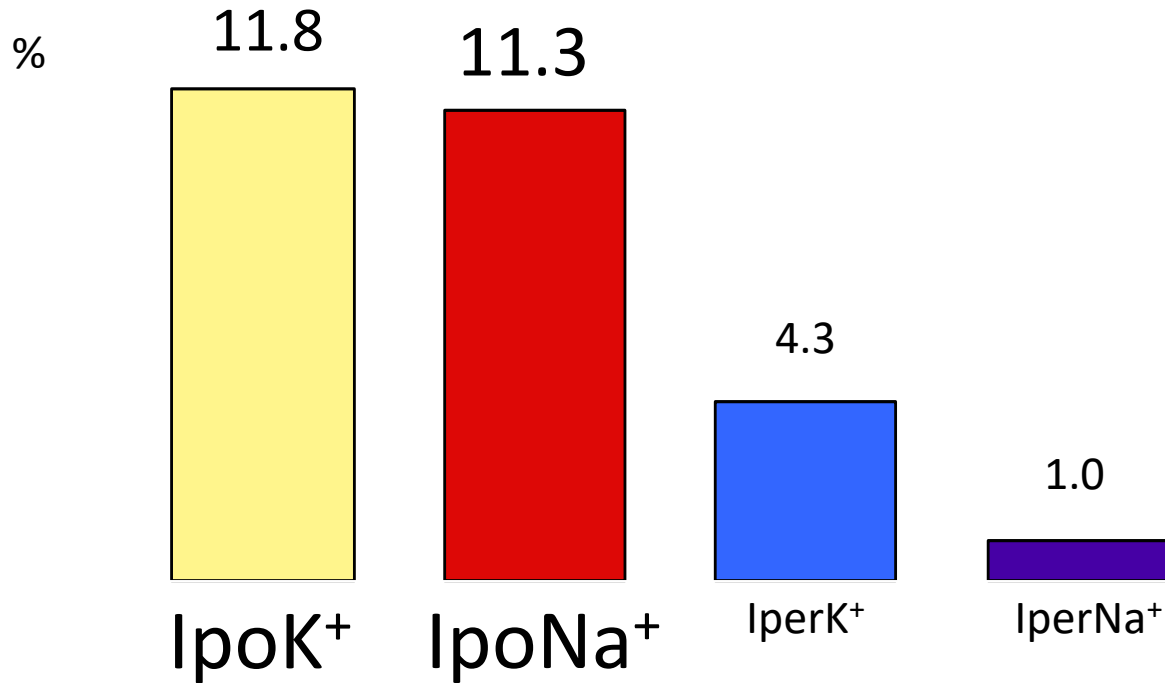
**Prof. Mauro Giordano**

Dir. Scuola di Specializzazione in Medicina d' Emergenza - Urgenza  
Seconda Università degli Studi di Napoli

NAPOLI 19\_11\_2016



# Prevalance of Electrolyte Disorders in Hospitalized Elderly



*Arief, DeFronzo*

*Fluid, Electrolyte and Acid-base Disorders, Churchill Livingstone 1995*

# General characteristics of patients with electrolyte imbalance admitted to emergency department

Arif Kadri Balci, Ozlem Koksal, Ataman Kose, Erol Armagan, Fatma Ozdemir, Taylan Inal, Nuran Oner

Faculty of Medicine, Department of Emergency Medicine, Uludag University, Bursa, Turkey

Corresponding Author: Ataman Kose, Email: ataberk76@yahoo.com.tr

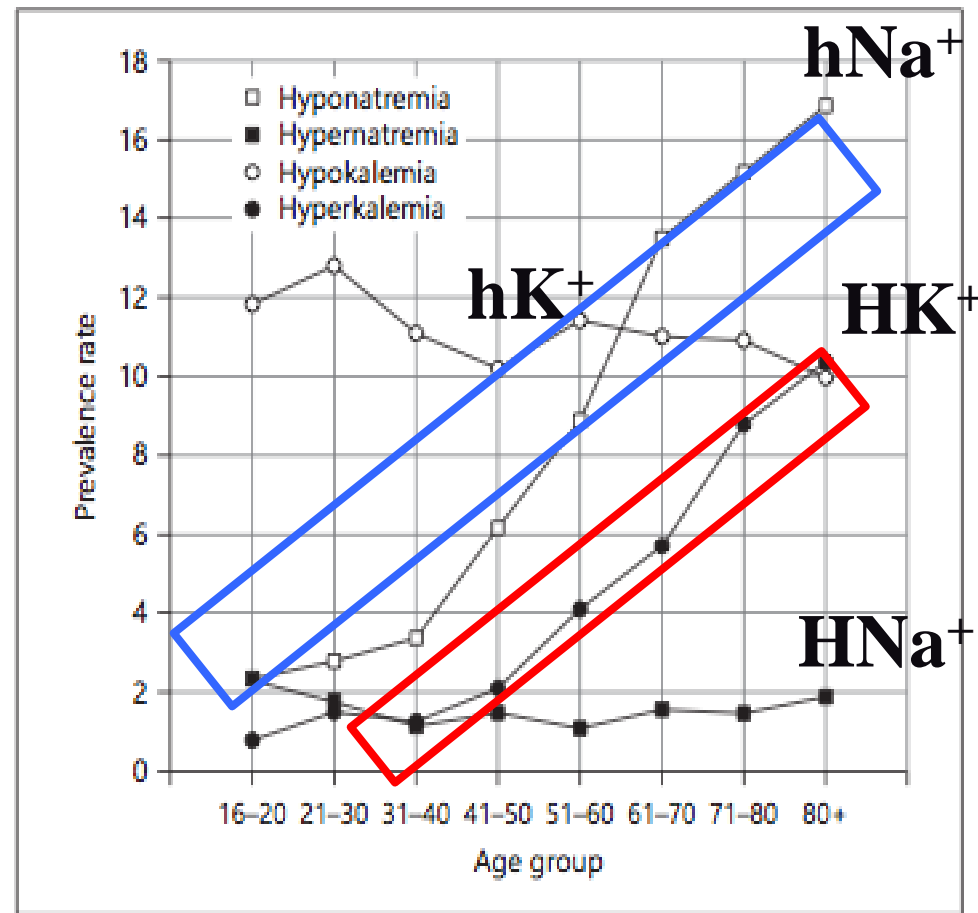
**Table 1.** Distribution of electrolyte imbalance

Electrolytes	Imbalance	Number of patients	%
Na <sup>+</sup>	Hyponatremia	600	60
	Hypernatremia	52	5
K <sup>+</sup>	Hypokalemia	152	15
	Hyperkalemia	80	8
Ca <sup>++</sup>	Hypocalcemia	512	51
	Hypercalcemia	38	4
Mg <sup>++</sup>	Hypomagnesemia	52	5
	Hypermagnesemia	10	1

## Age-Related Variety in Electrolyte Levels and Prevalence of Dysnatremias and Dyskalemias in Patients Presenting to the Emergency Department

Gregor Lindner<sup>a</sup> Carmen A. Pfortmüller<sup>a</sup> Alexander B. Leichtle<sup>b</sup>  
Georg M. Fiedler<sup>b</sup> Aristomenis K. Exadaktylos<sup>a</sup>

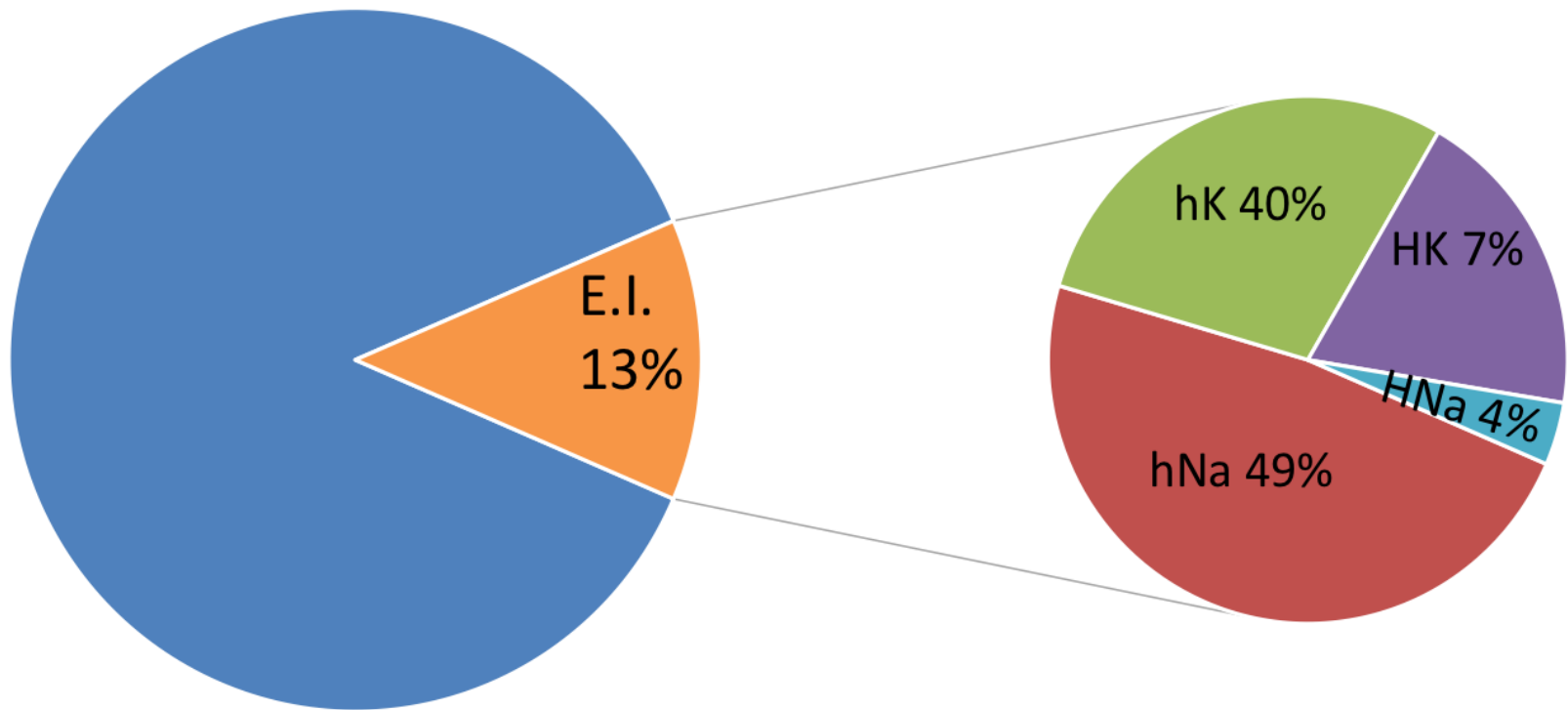
<sup>a</sup>Department of Emergency Medicine and <sup>b</sup>Center for Laboratory Medicine, Inselspital, University Hospital Bern, Bern, Switzerland



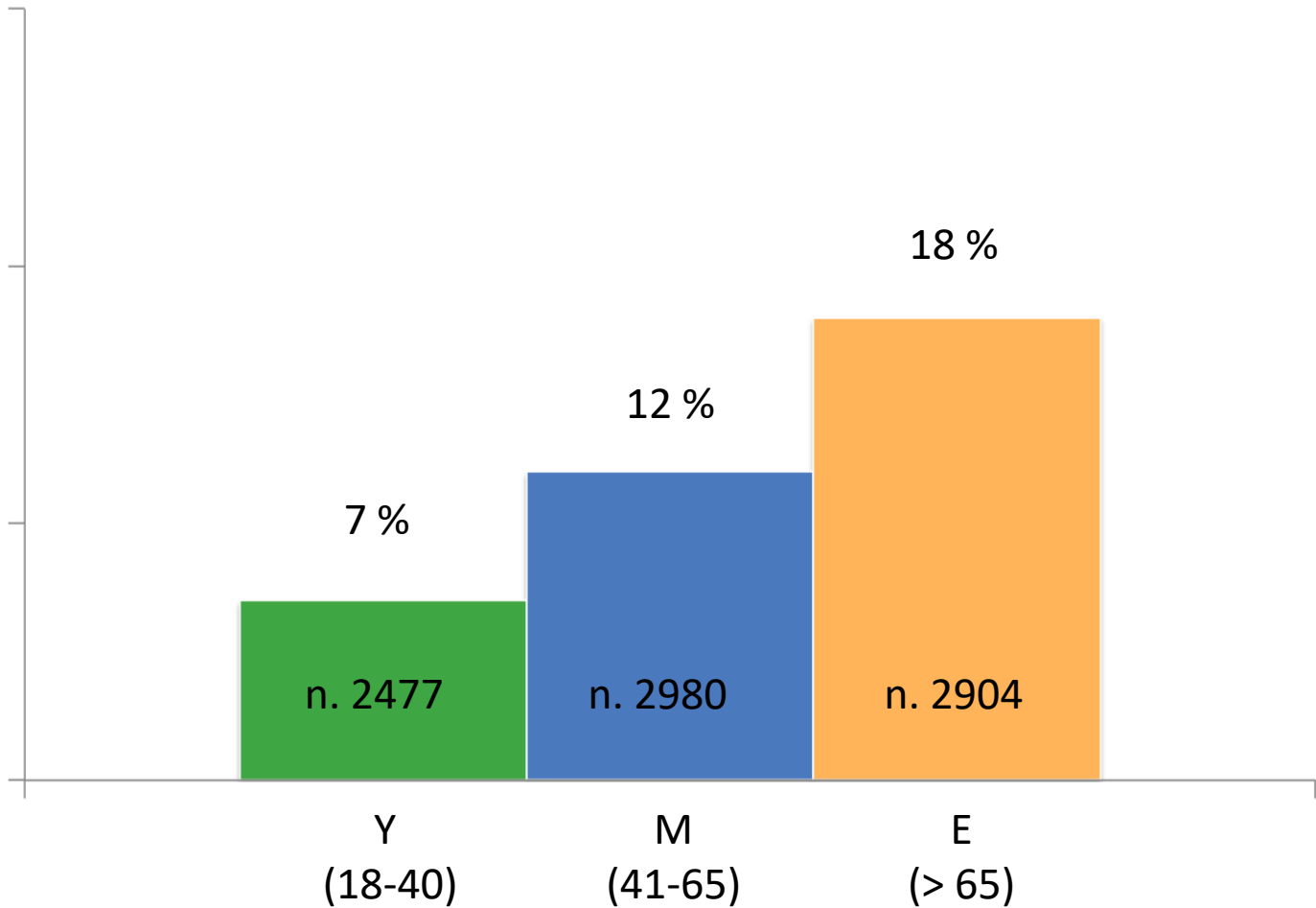
**Fig. 1.** Prevalence rates of dysnatremias and dyskalemias stratified for age group.

# PREVALANCE OF ELECTROLYTE IMBALANCE IN E.D.

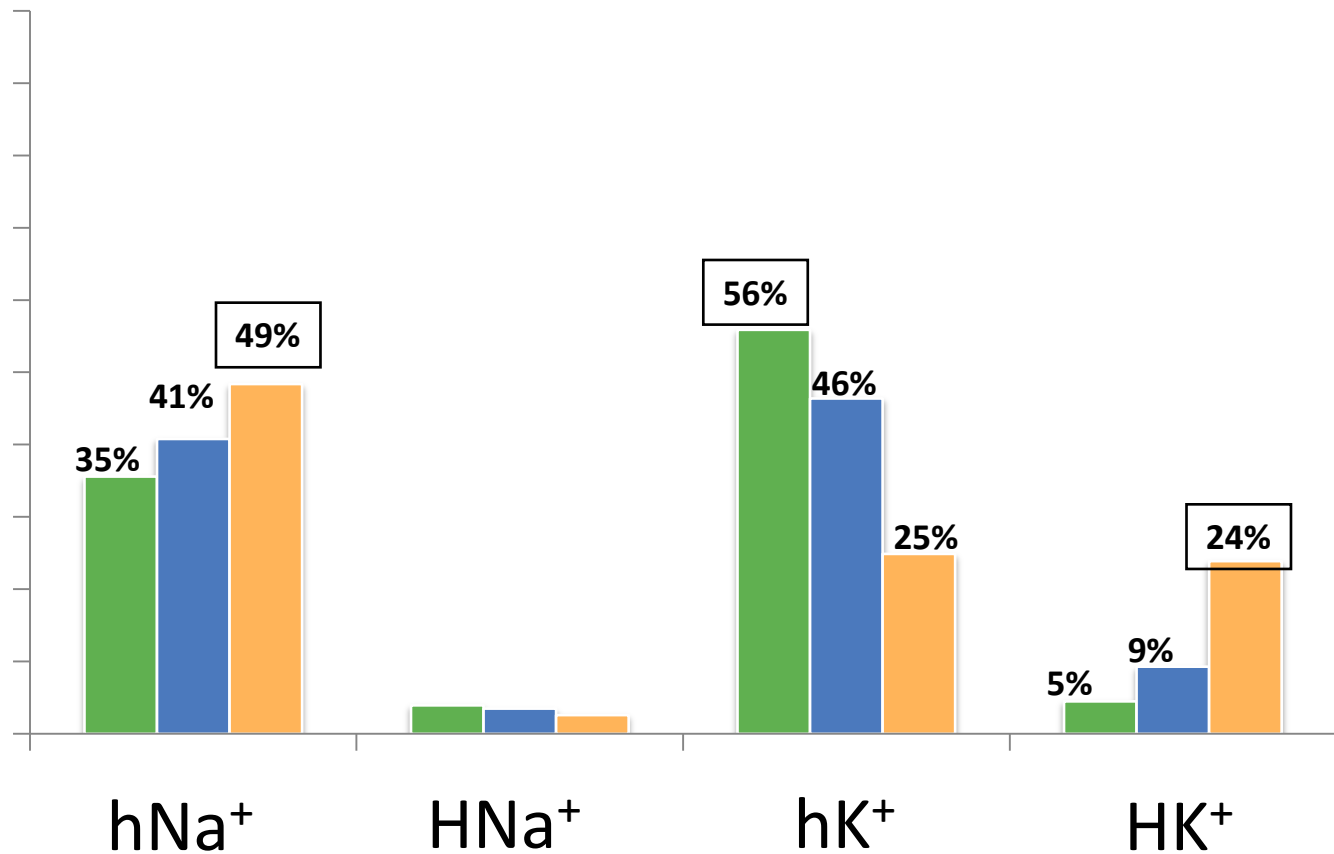
n = 8631 (2015)



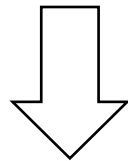
# AGE RELATED PREVALANCE OF ELECTROLYTE IMBALANCE IN E.D.



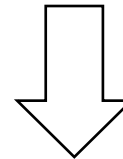
# AGE RELATED PREVALANCE OF ELECTROLYTE IMBALANCE IN E.D.



# OMEOSTASI DEL SODIO



98 %



2 %



# Relationship between vascular disease and age-associated changes in the human kidney

BERTRAM L. KASISKE

Department of Medicine, Hennepin County Medical Center, University of Minnesota, Minneapolis, Minnesota, USA

— (Group I): **Mild** Atherosclerosis

- - - (Group II): **Severe** Atherosclerosis

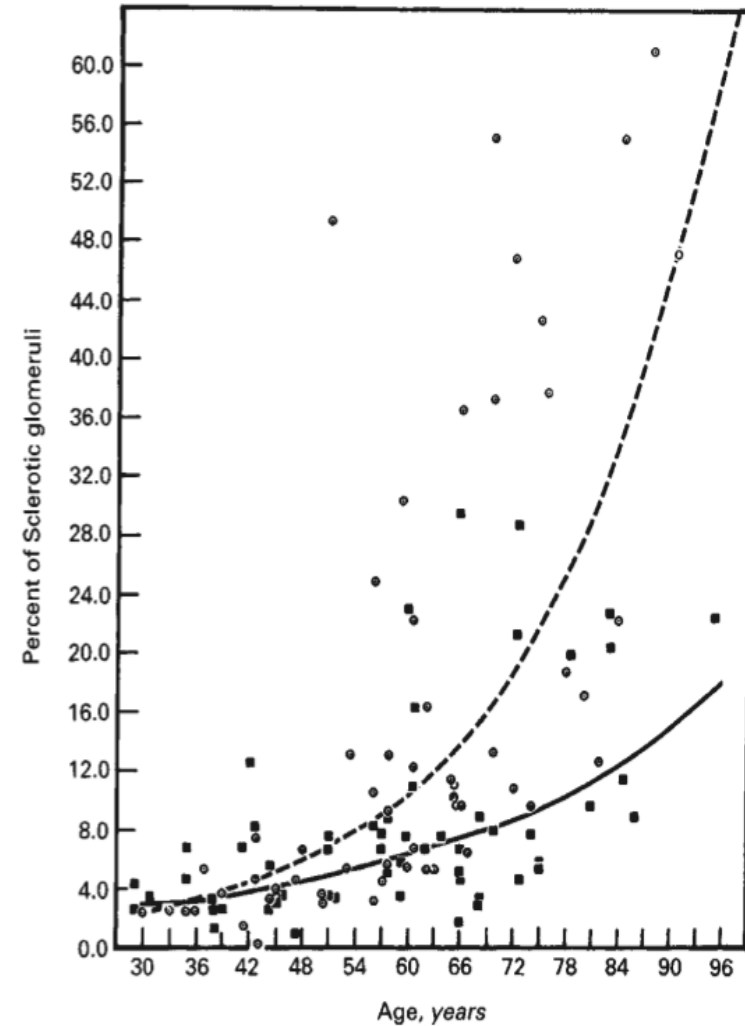


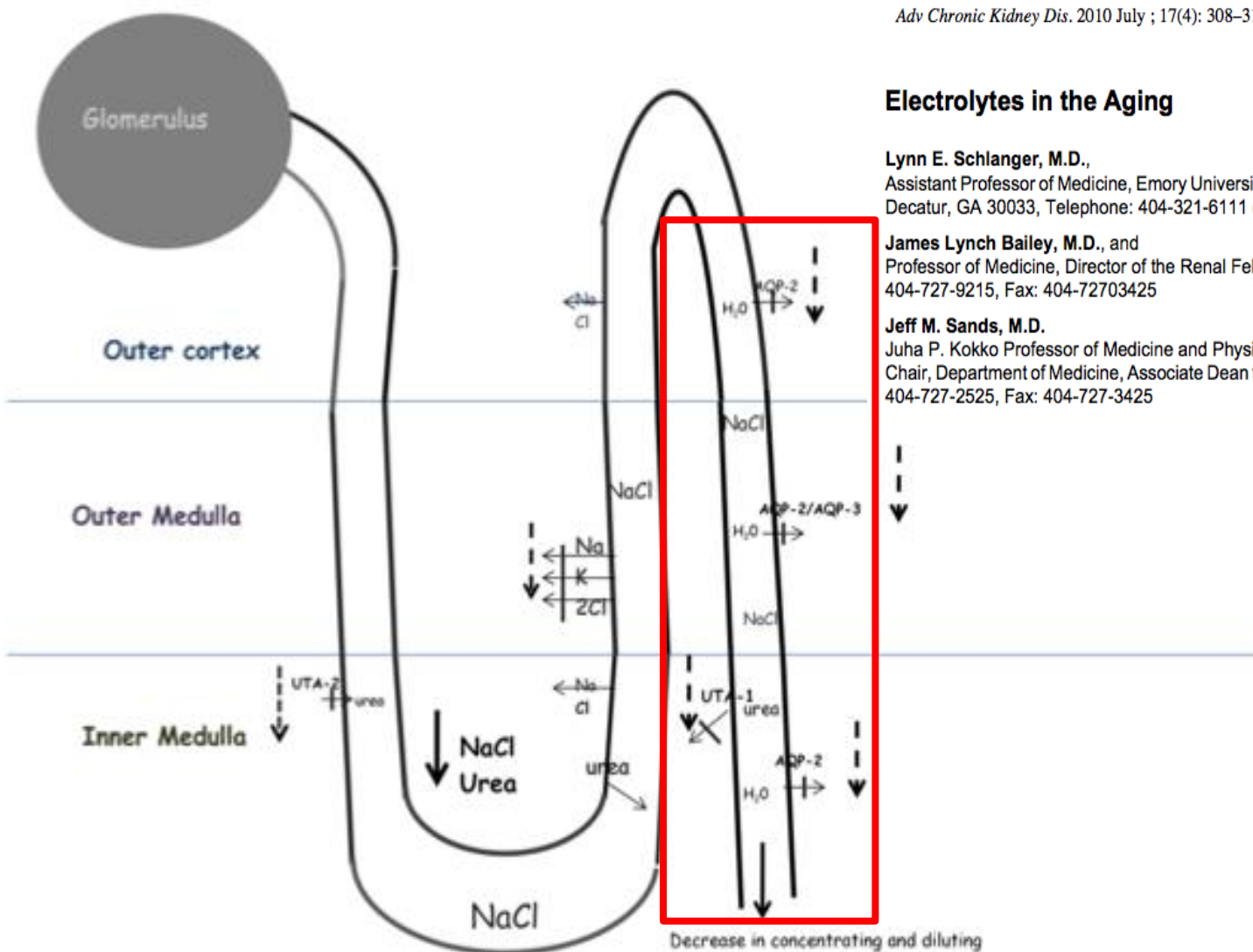
Fig. 1. The influence of systemic atherosclerosis on the relationship between age and glomerulosclerosis. Least squares best fit lines for logarithm of glomerulosclerosis (%) vs. age for group I (solid points and solid line) and group II (open circles and dashed line).

## The Influence of Age on the Renal Response to Water Deprivation in Man

JOHN W. ROWE, NATHAN W. SHOCK and RALPH A. DEFONZO

Clinical Physiology Branch, Gerontology Research Center, National Institute  
on Aging, National Institutes of Health, Baltimore, Md.

	Mean age, years	Urine osmolality, mosm/kg <sup>1</sup>		Urine flow, ml/min <sup>1</sup>	
		period 1	period 3	period 1	period 3
Young, 20-39 (n = 31)	33	969 ±41	1,109 ±22	1.02 ±0.10	0.49 ±0.03
Middle, 40-59 (n = 48)	49	949 ±39	1,051 ±19	0.99 ±0.10	0.63 ±0.03
Old, 60-79 (n = 18)	68	852 ±64	882 ±49	1.05 ±0.15	1.03 ±0.13



## Electrolytes in the Aging

Lynn E. Schlanger, M.D.,

Assistant Professor of Medicine, Emory University/VAMC at Atlanta, Address 1670 Clairmont Road, Decatur, GA 30033, Telephone: 404-321-6111 ext 7070, Fax: 404-235-3049

James Lynch Bailey, M.D., and

Professor of Medicine, Director of the Renal Fellowship Program, Emory University, Telephone: 404-727-9215, Fax: 404-72703425

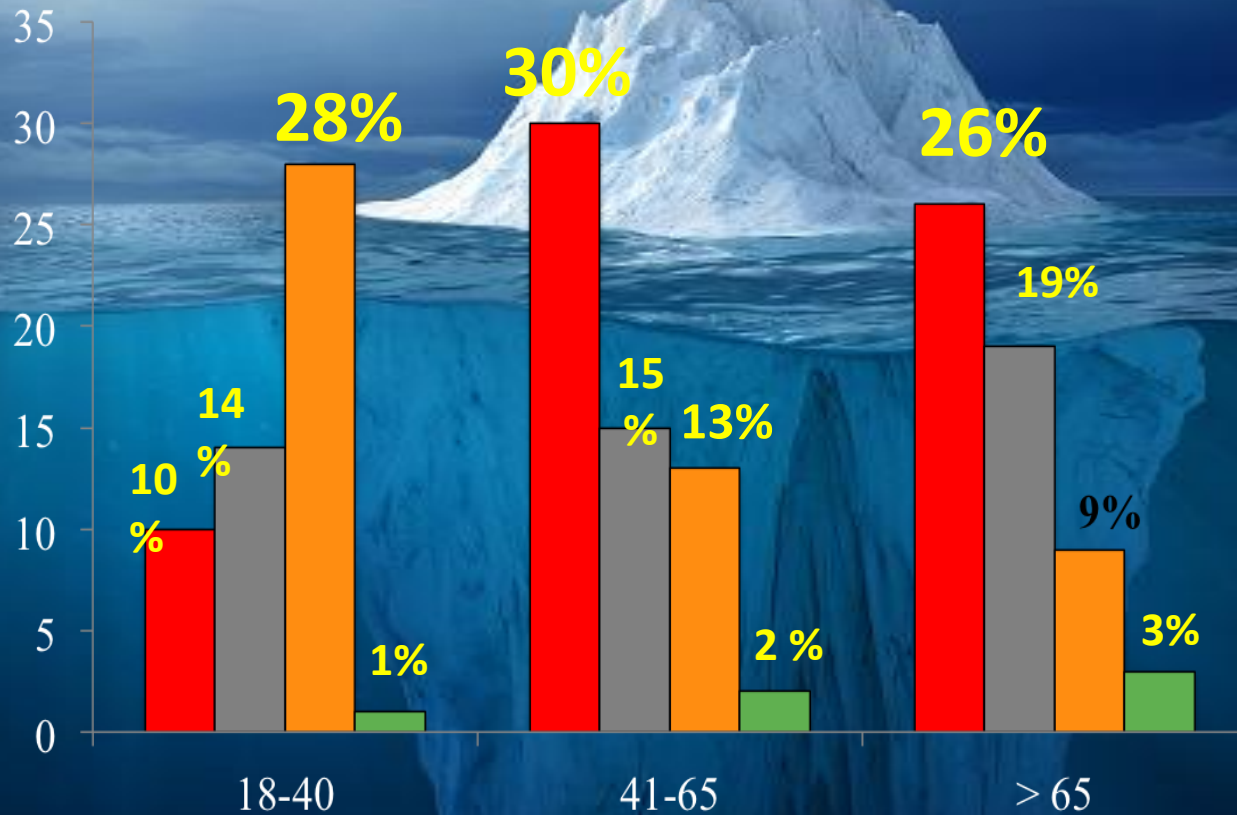
Jeff M. Sands, M.D.

Juha P. Kokko Professor of Medicine and Physiology, Director, Renal Division, Executive Vice-Chair, Department of Medicine, Associate Dean for Clinical and Translational Research, Telephone: 404-727-2525, Fax: 404-727-3425

Fig 2.

The putative changes in the renal transport system in the elderly are shown. In animal studies there is a decrease in the abundance of AQP2, AQP3, NKCC2/BSC1 and UT-A1, A2, A3. The arrows represent the transporters and water channels known to be downregulated in animal studies. These changes may be present in the elderly population affecting the diluting and concentrating capacity.

# PATOLOGY ASSOCIATED WITH ELECTROLYTE DISORDERS IN E.D.



**CV** **LD** **GI** **EI**

## VII Joint National Committee on Prevention, Detection, Evaluation and Treatment of High Blood Pressure: JAMA 03

### Lifestyle Modifications in the Management of Hypertension<sup>†</sup>

<b>Modification</b>	<b>Recommendation</b>	<b>Approximate systolic BP reduction, range*</b>
Weight reduction	Maintain normal body weight (BMI, 18.5 to 24.9 kg/m <sup>2</sup> )	5-20 mmHg per 10-kg weight loss
Adopt DASH eating plan	Consume a diet rich in fruits, vegetables, and low-fat dairy products with a reduced content of saturated and total fat	8 to 14 mmHg
Dietary sodium reduction	Reduce dietary sodium intake to no more than 100 meq/day (2.4 g sodium or 6 g sodium chloride)	2 to 8 mmHg
Physical activity	Engage in regular aerobic physical activity such as brisk walking (at least 30 minutes per day, most days of the week)	4 to 9 mmHg
Moderation of alcohol consumption	Limit consumption to no more than 2 drinks per day in most men and no more than 1 drink per day in women and lighter-weight persons	2 to 4 mmHg

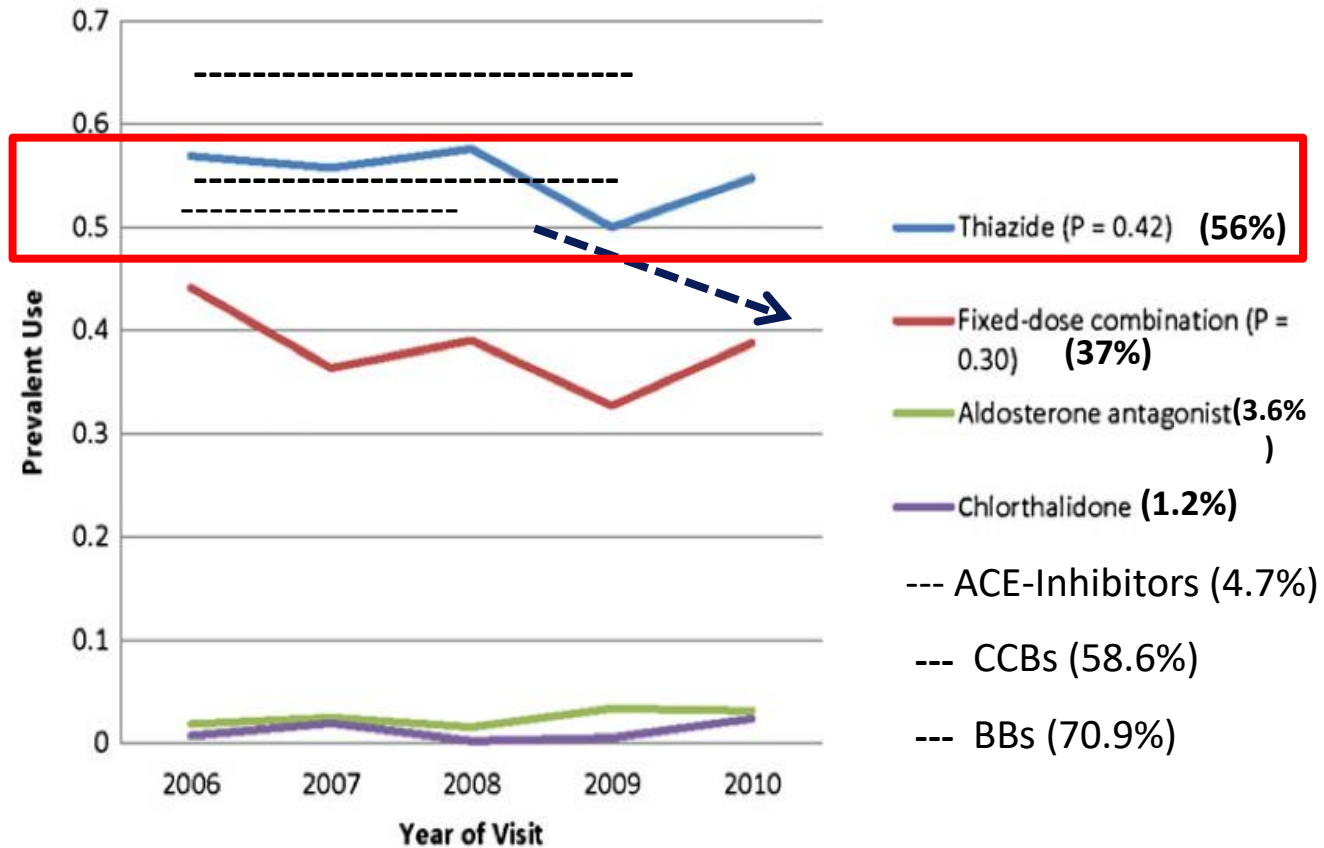
# Iponatremia ed Ipertensione

Vi è una correlazione?

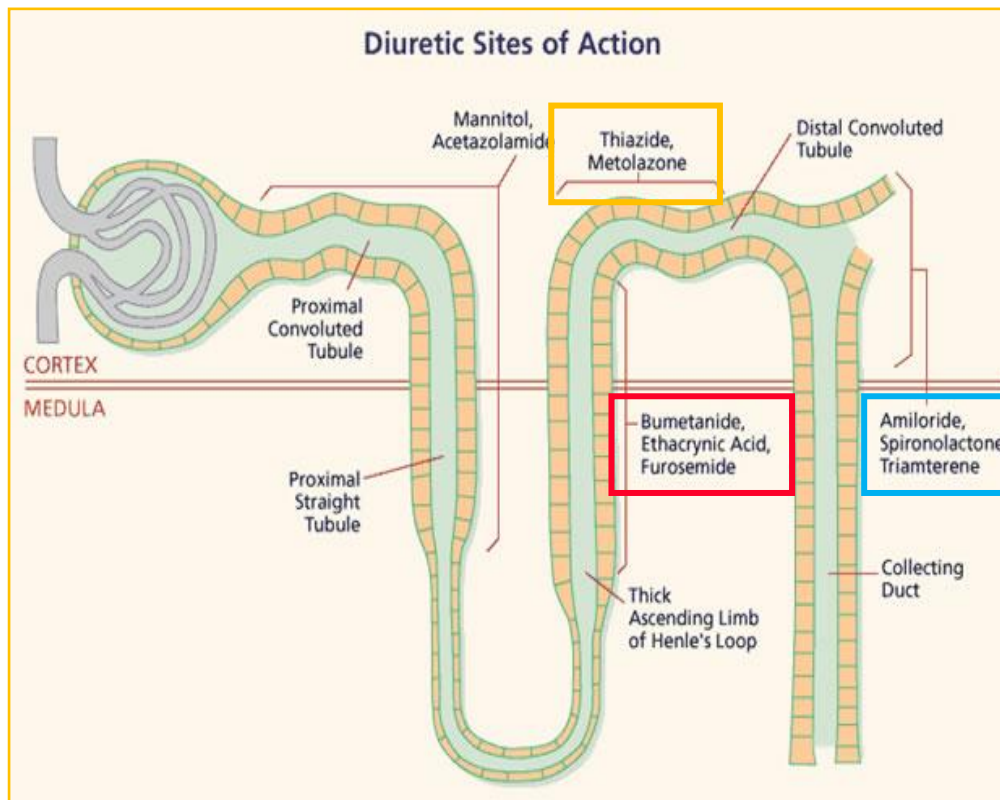
In Italia si stima che circa 12 milioni di abitanti sono ipertesi, ben il 20% della popolazione!



# USE OF DIURETICS IN USA







<b>Nephron Segment</b>	<b>Diuretic</b>	<b>Action</b>
Loop of Henle	Furosemide, Torasemide, Bumetanide	Na <sup>+</sup> - K <sup>+</sup> 2Cl <sup>-</sup> cotransport
Distal Tubule	Thiazides, Chlorthalidone, Indapamide	Na <sup>+</sup> - Cl <sup>-</sup> cotransport
Collecting Ducts	Spirolactone, Amiloride, Triamterene	Na <sup>+</sup> channels



# Water Load and Thiazide-Induced Hyponatremia

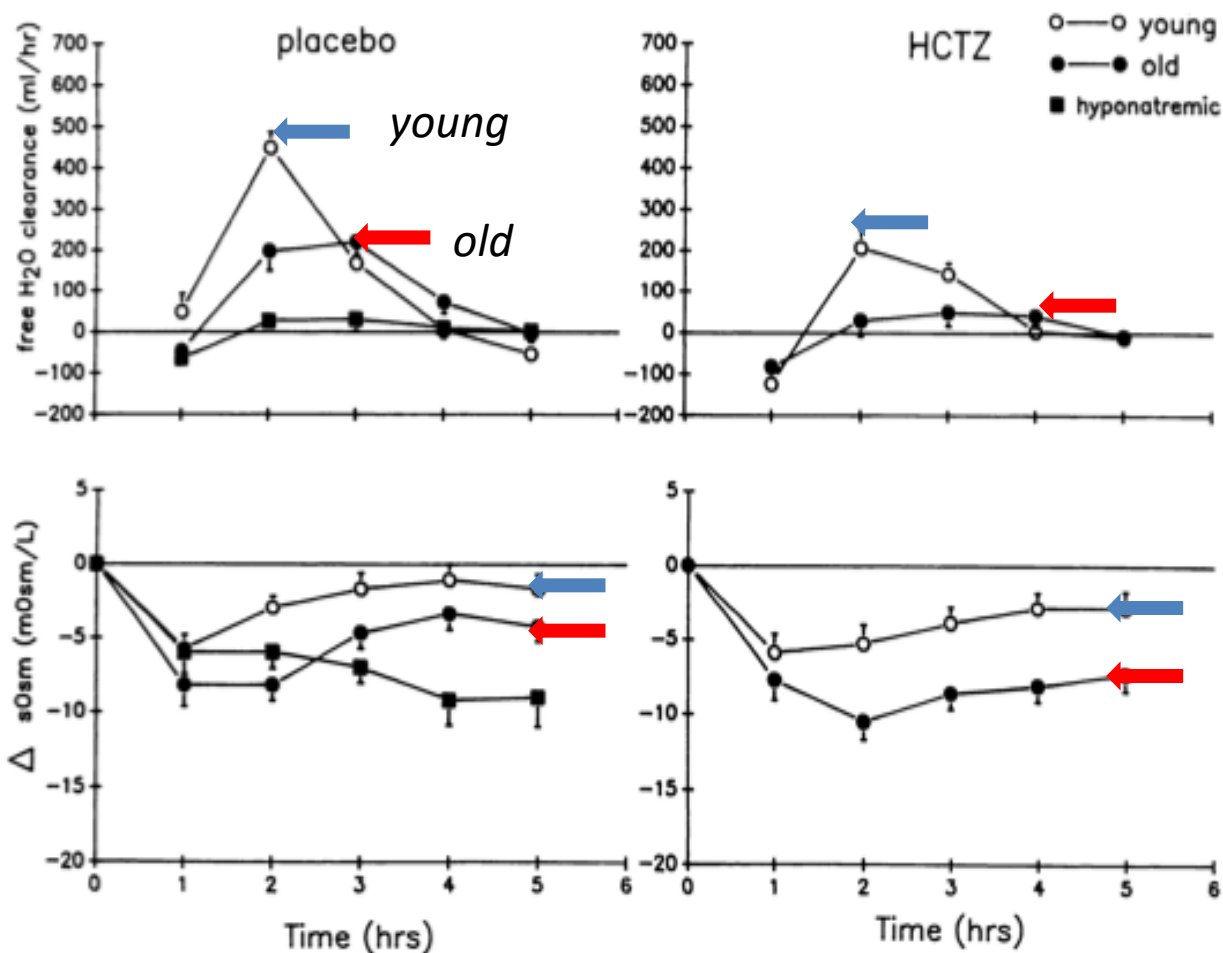
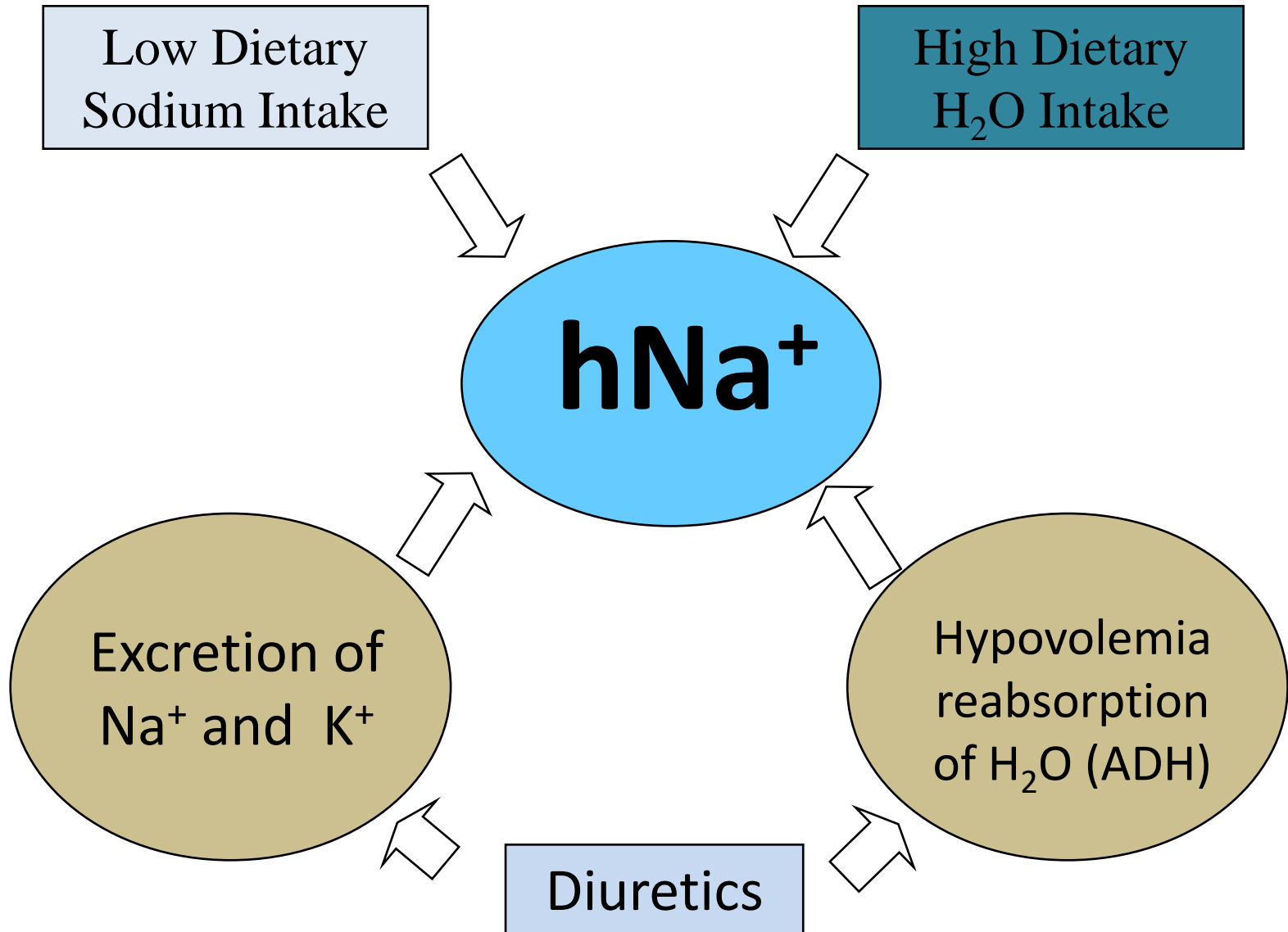
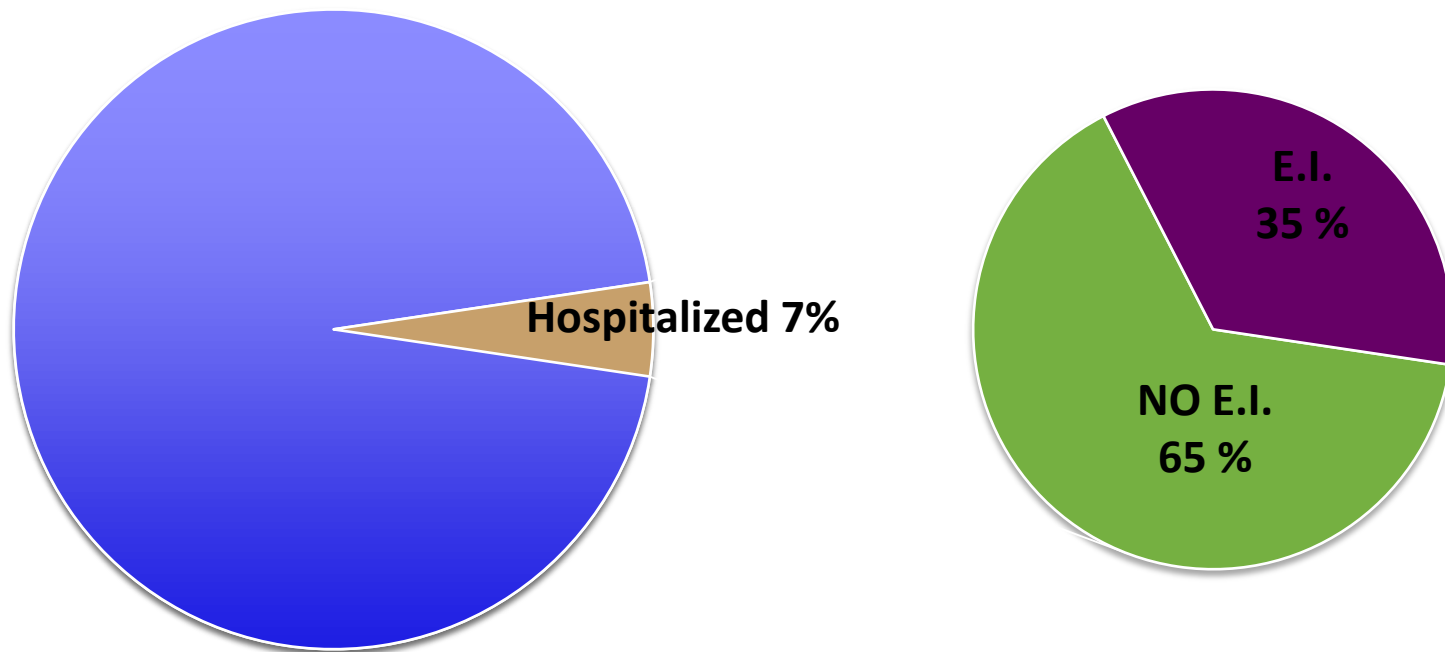


Figure 1. Free water clearance (CH<sub>2</sub>O) and change in serum osmolality (ΔsOsm, mosm/kg H<sub>2</sub>O) after a water load with placebo versus HCTZ in young, old, and old with a prior history of thiazide-induced hyponatremia. CH<sub>2</sub>O and decline in sOsm were significantly lower in the old than in the young ( $P < 0.05$ , ANOVA). This difference was magnified after the use of HCTZ. Those with a history of hyponatremia had lower CH<sub>2</sub>O and decline in sOsm than did the healthy elderly ( $P < 0.05$ , ANOVA).

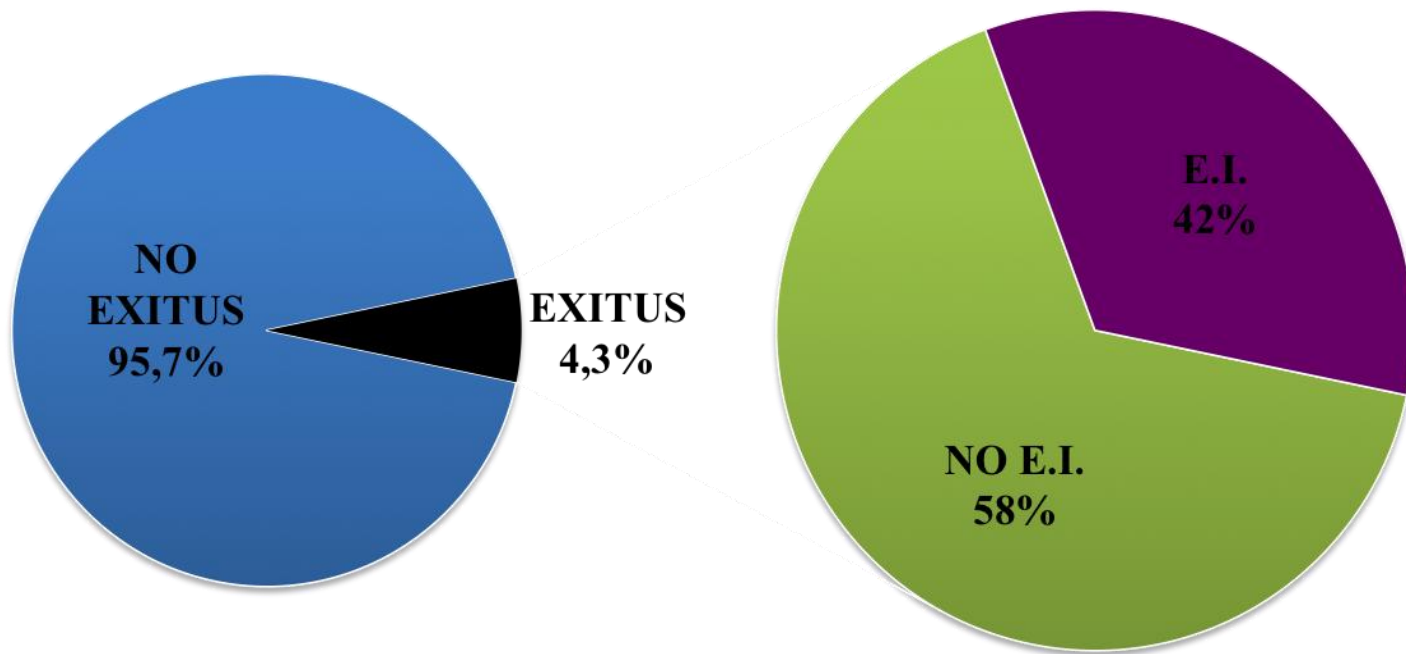
# GENESIS OF HYPONATREMIA



# PREVALANCE OF ELECTROLYTE IMBALANCE IN HOSPITALIZED PATIENTS



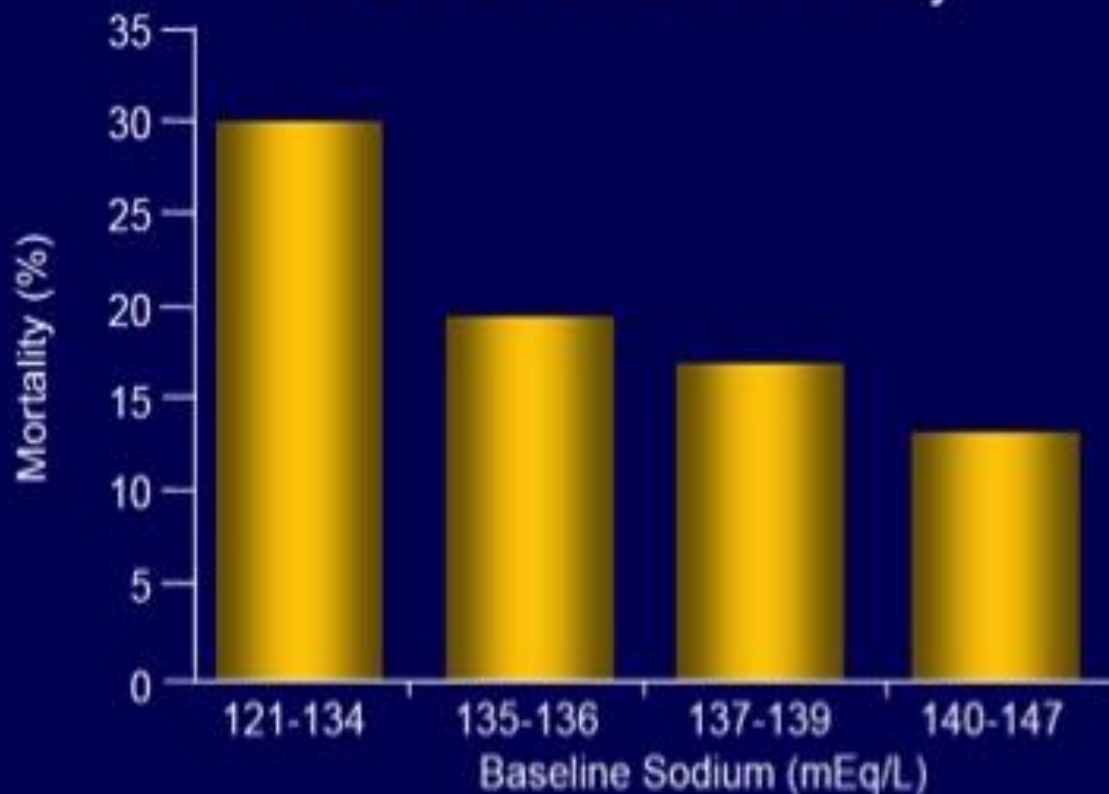
# PREVALANCE OF ELECTROLYTE IMBALANCE IN EXITUS



# Correlation Between Hyponatremia and Mortality Risk in Heart Failure Patients

---

ESCAPE - 6 month mortality



ESCAPE=Evaluation Study of Congestive Heart Failure and Pulmonary Artery Catheterization Effectiveness

Gheorghiade M, et al. Arch Intern Med. 2007;167:1998-2005.



Original article

## Hyponatremia is an independent predictor of adverse clinical outcomes in hospitalized patients due to worsening heart failure

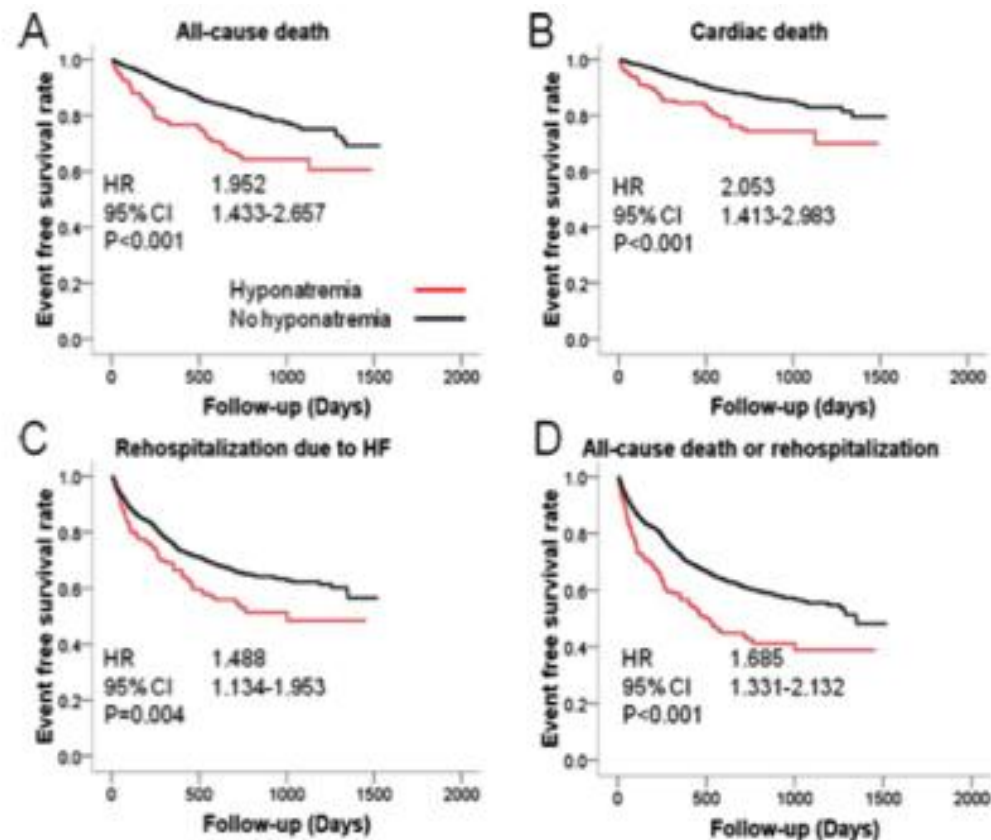


Sanae Hamaguchi (MD, PhD)<sup>a,b,1</sup>, Shintaro Kinugawa (MD, PhD)<sup>a,1</sup>,  
Miyuki Tsuchihashi-Makaya (RN, PhD)<sup>c,1</sup>, Shouji Matsushima (MD, PhD)<sup>a,1</sup>,  
Mamoru Sakakibara (MD, PhD)<sup>a,1</sup>, Naoki Ishimori (MD, PhD)<sup>a,1</sup>,  
Daisuke Goto (MD, PhD)<sup>a,1</sup>, Hiroyuki Tsutsui (MD, PhD, FJCC)<sup>a,\*</sup>

<sup>a</sup> Department of Cardiovascular Medicine, Hokkaido University Graduate School of Medicine, Sapporo 060-8638, Japan

<sup>b</sup> Department of Cardiovascular Medicine, Social Welfare Corporation Hokkaido Social Work Association Obihiro Hospital, Obihiro 080-0805, Japan

<sup>c</sup> School of Nursing, Kitasato University, Sagamihara 252-0329, Japan

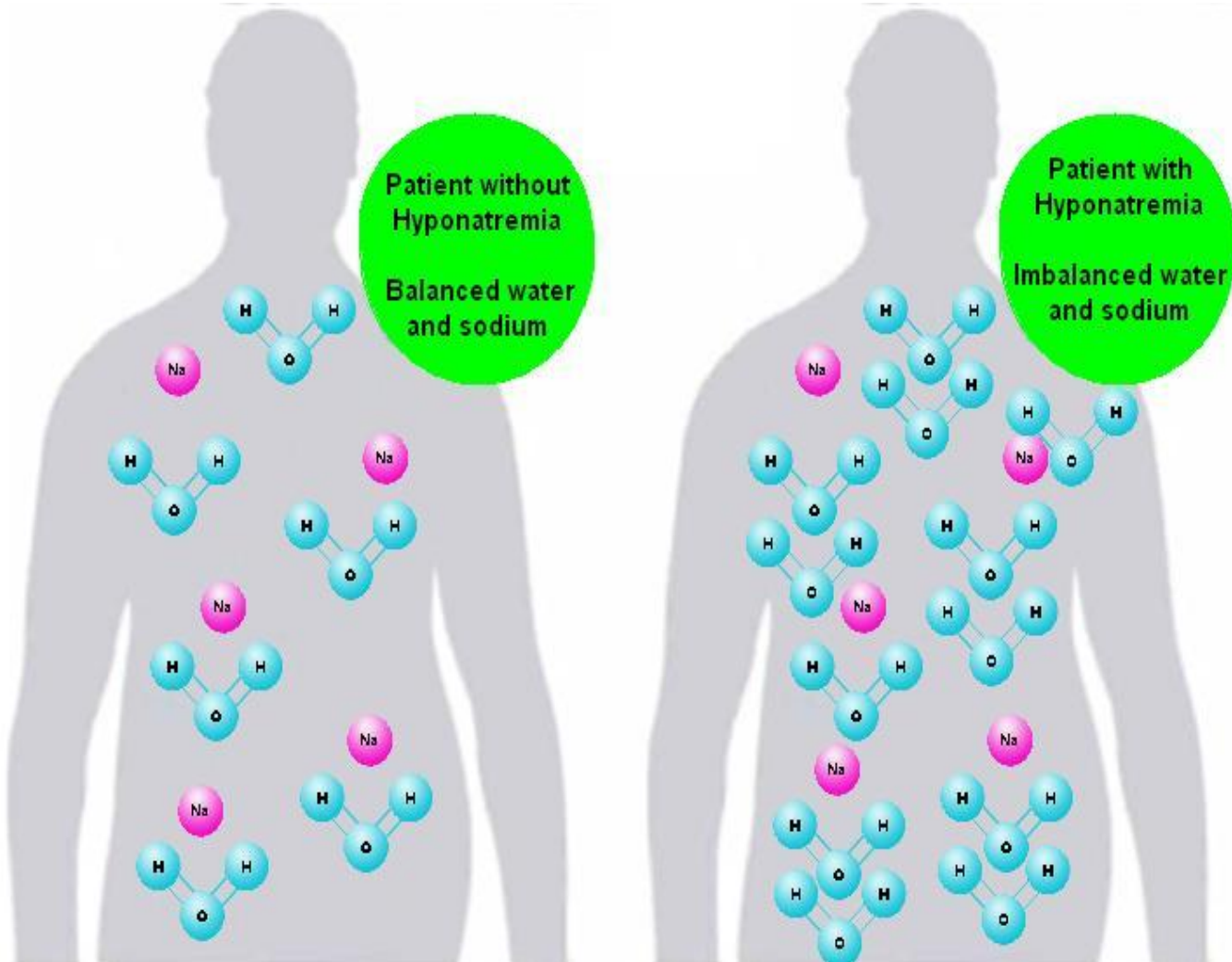


**Fig. 2.** Kaplan–Meier survival curves free from all-cause death (A), cardiac death (B), rehospitalization due to worsening heart failure (HF) (C), and all-cause death or rehospitalization (D) according to the presence or absence of hyponatremia.

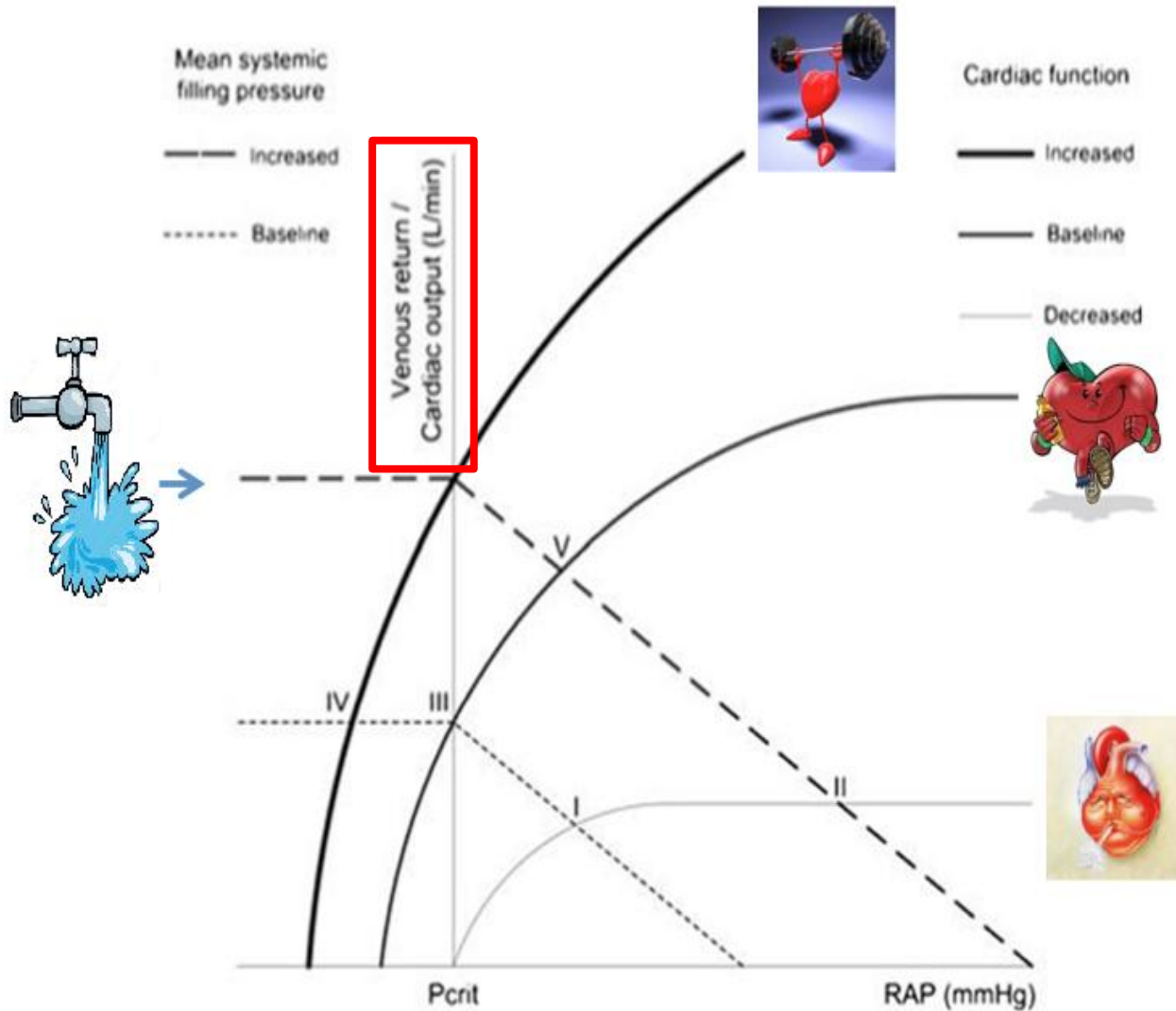
# Hyponatremia

< 60% water

> 60% water

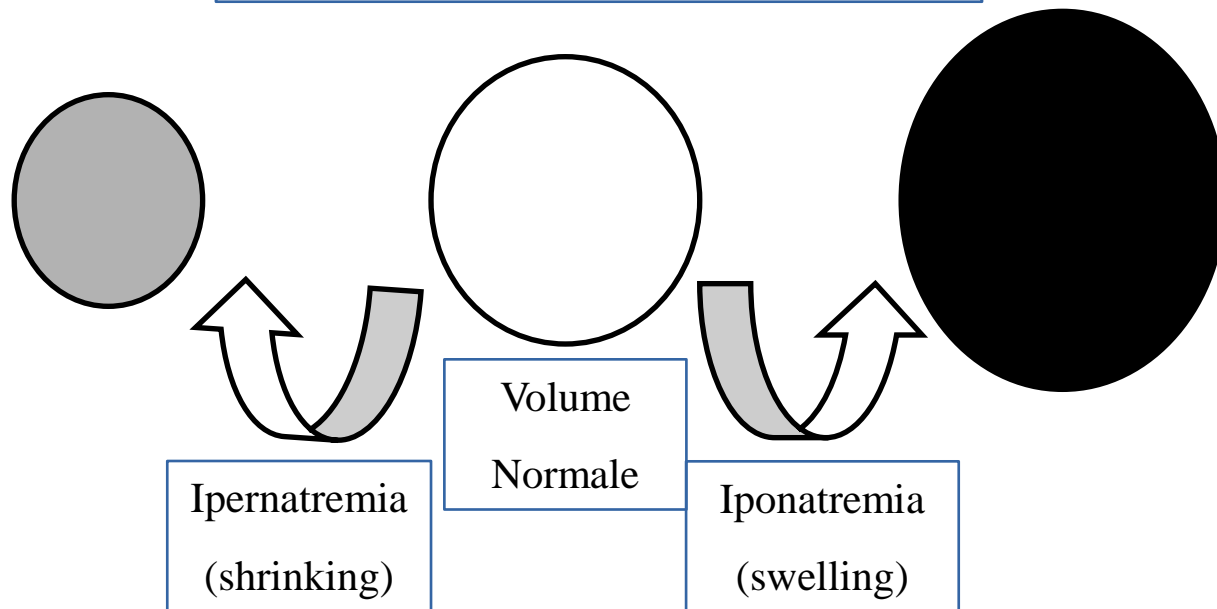


# Cardiac Function and Water

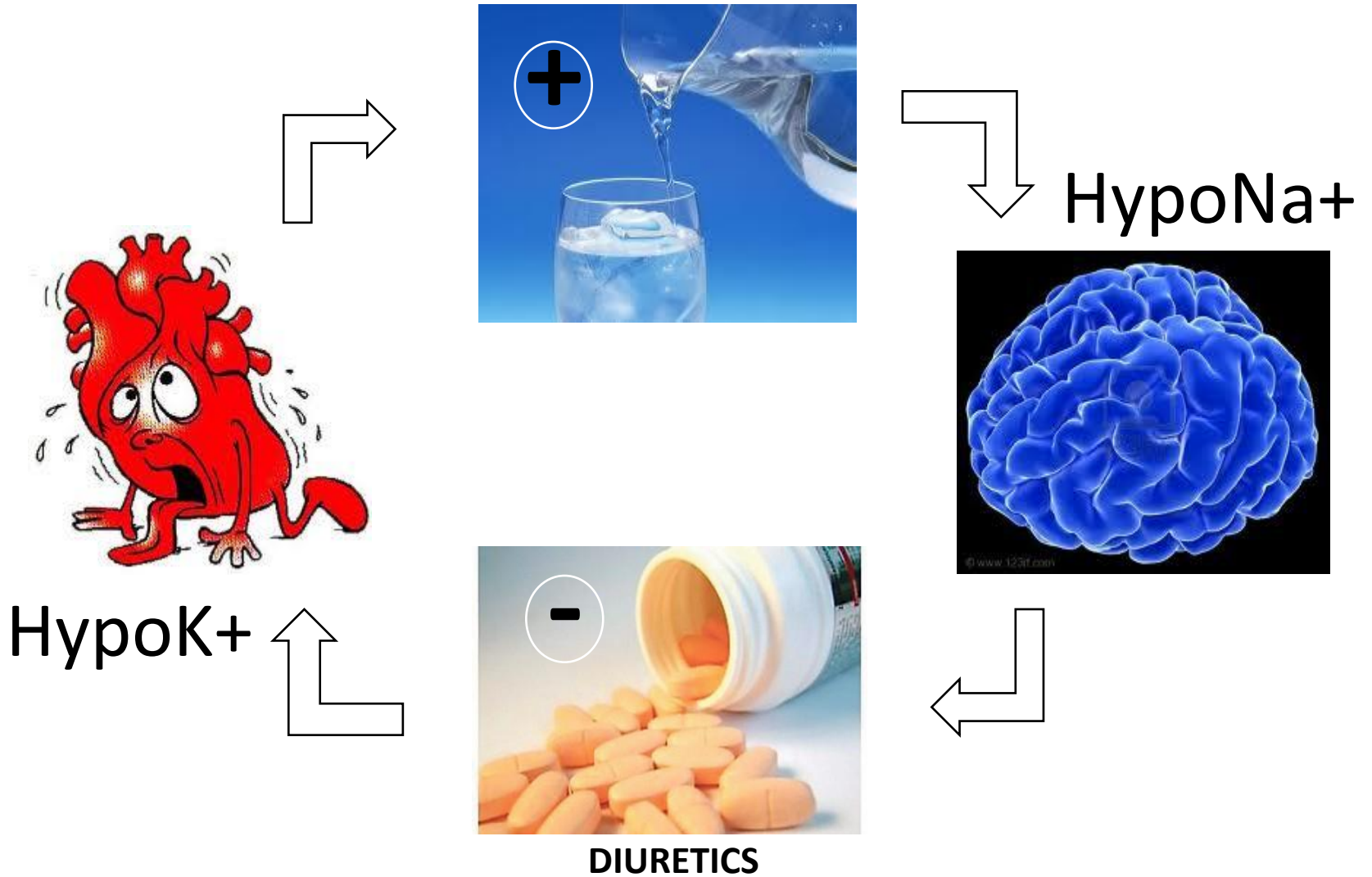




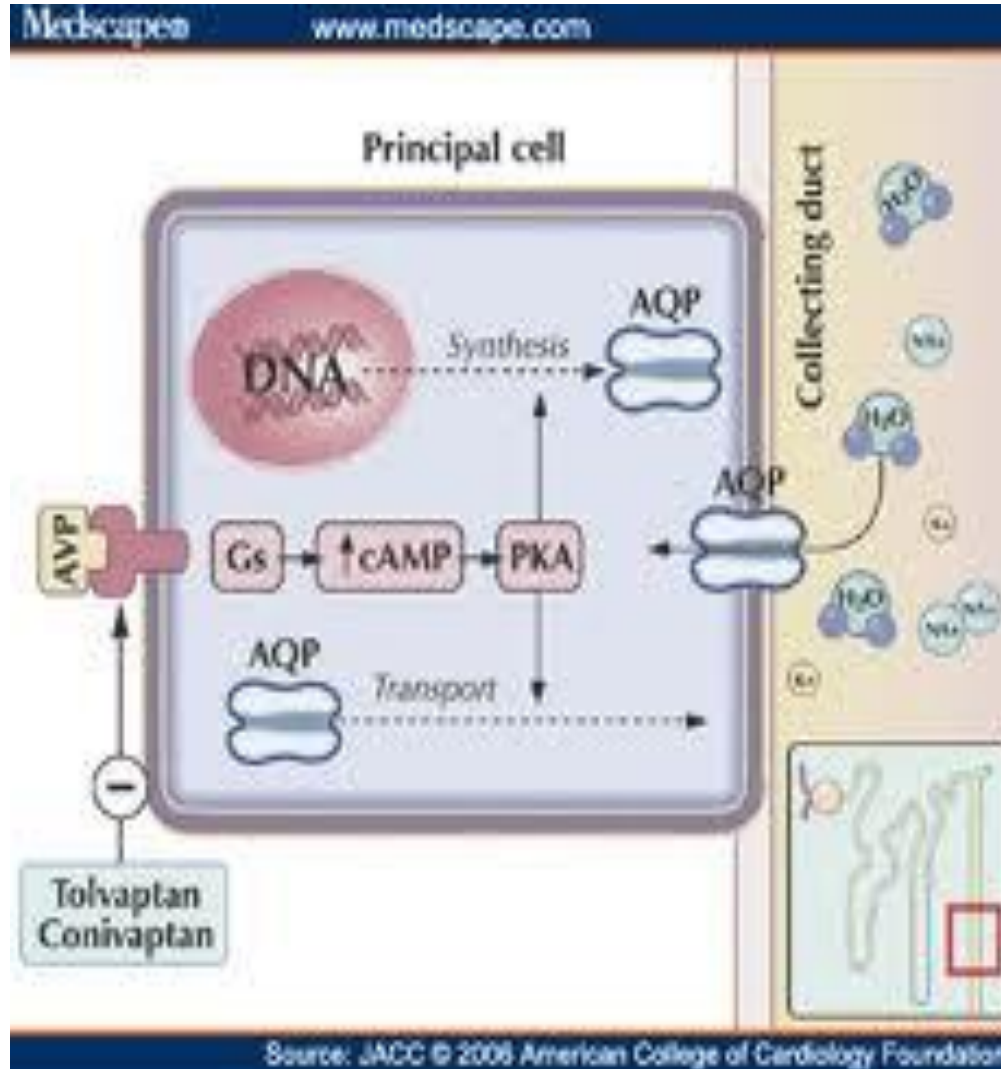
# Stress Osmotico della Cellula Cerebrale



# The Vicious Cycle of Water



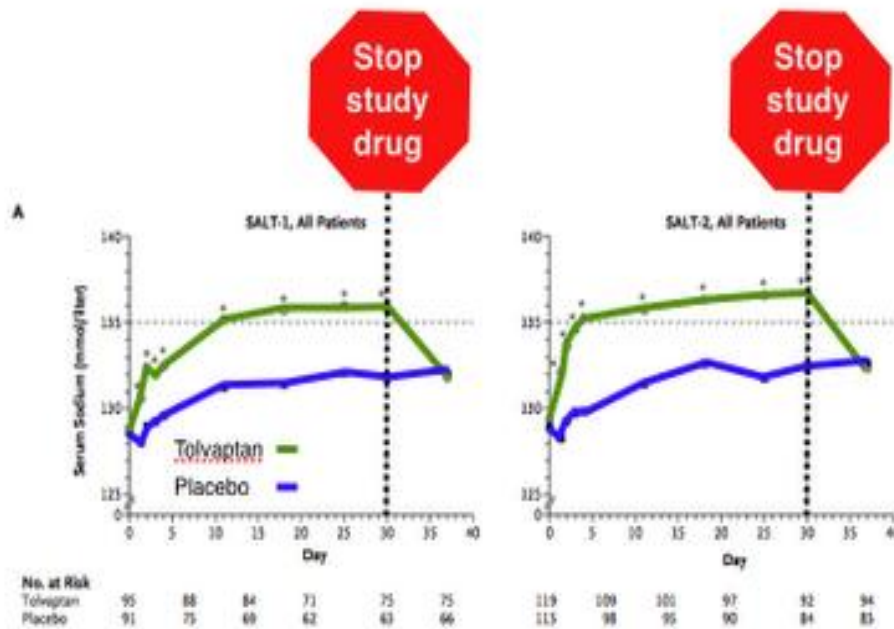
# UNA NUOVA CLASSE DI DIURETICI: I VAPTANI



# Tolvaptan a Selective Oral Vasopressin V2-Receptor Antagonist for Hyponatremia

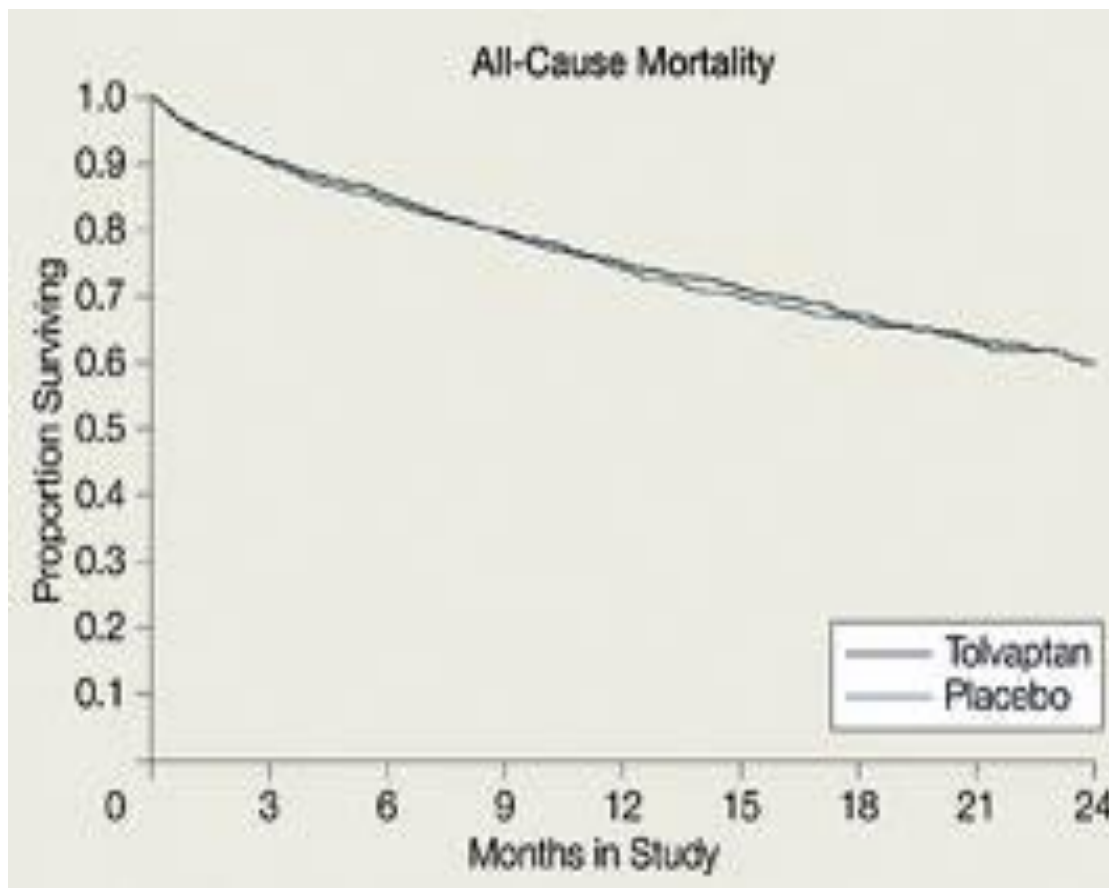
Robert W. Schrier, M.D for the SALT Investigators

## Tolvaptan 15, 30, 60 mg orally OD



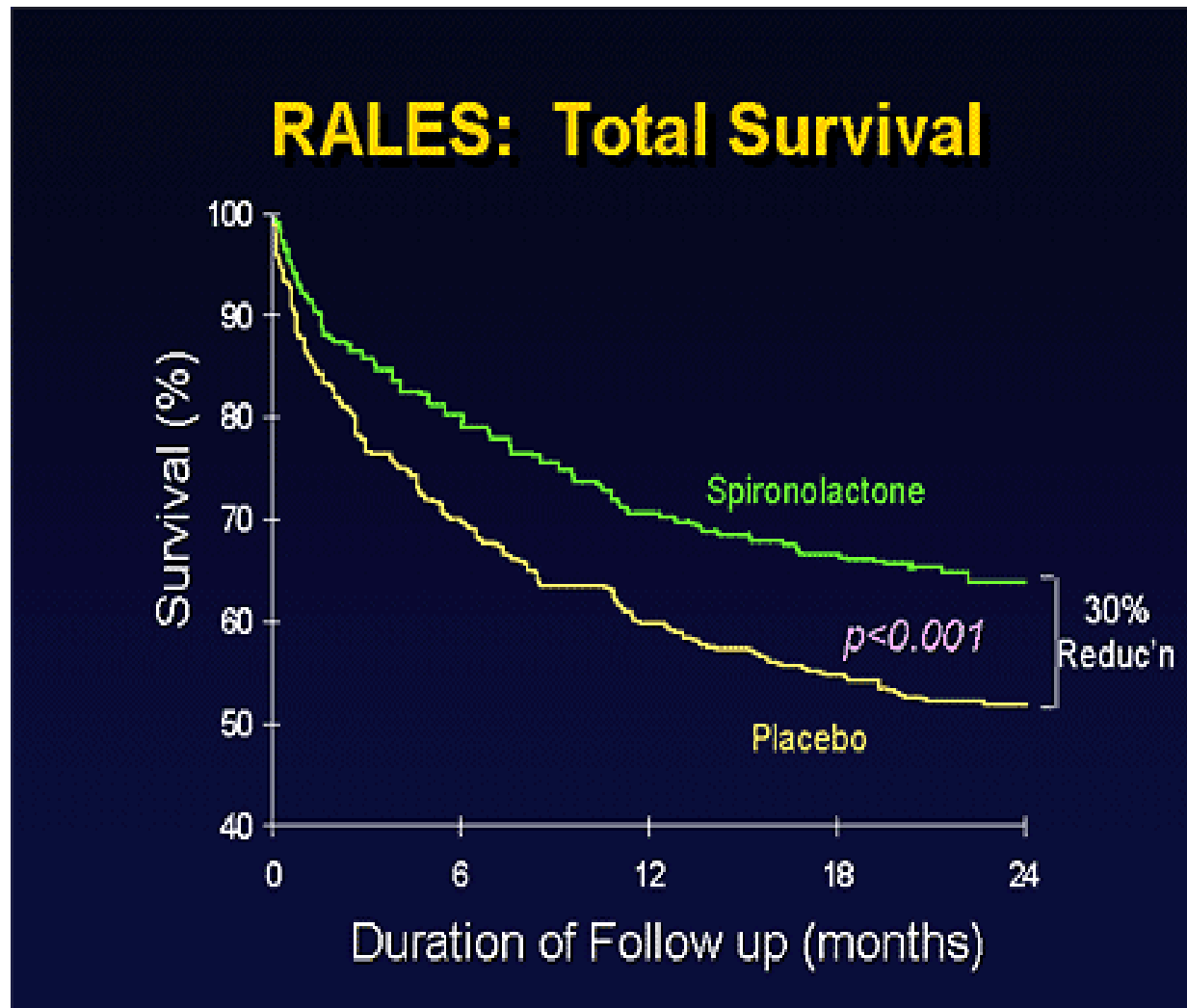
# Effects of Oral Tolvaptan in Patients Hospitalized for Worsening Heart Failure

The EVEREST Outcome Trial

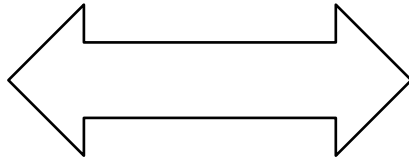


# Spironolactone & HF

The Randomized Aldactone Evaluation Study Trial



# L'acqua vita o morte ?



# IPONATREMIA

## IPO-EUVOLEMICA

## IPERVOLEMICA

Deficit totale di  $\text{Na}^+$  > Deficit totale di  $\text{H}_2\text{O}$

Eccesso totale di  $\text{H}_2\text{O}$  > Eccesso totale di  $\text{Na}^+$

### Deplezione ECF

### Eccesso ECF

#### Perdita renale

- Eccesso di diuretici
- Deficit mineralcorticoidi
- "Salt-losing Nephritis"

#### Perdita extrarenale

- Vomito
- Diarrea
- Terzo spazio
- Pancreatite
- Peritonite
- Trauma muscolare

Sodiuria  
> 20  
mmol/l

Sodiuria  
< 10  
mmol/l

Soluzione salina  
isotonica

- Scompenso cardiaco
- Cirrosi
- Sindrome nefrosica

Sodiuria  
< 10  
mmol/l

IR  
acuta e  
cronica  
SIADH

Sodiuria  
> 20  
mmol/l

Soluzione Salina  
Ipertonica



# IPONATREMIA + SETE

## IPO-EUVOLEMICA

Deficit totale di  $\text{Na}^+$  > Deficit totale di  $\text{H}_2\text{O}$

Deplezione ECF

### Perdita renale

- Eccesso di diuretici
- Deficit mineralcorticoidi
- "Salt-losing Nephritis"

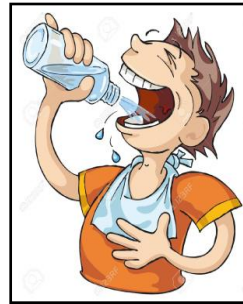
Sodiuria  
> 20  
mmol/l

Soluzione salina  
isotonica

### Perdita extrarenale

- Vomito
- Diarrea
- Terzo spazio
- Pancreatite
- Peritonite
- Trauma muscolare

Sodiuria  
< 10  
mmol/l



**H<sub>2</sub>O** →

## IPERVOLEMICA

Eccesso totale di  $\text{H}_2\text{O}$  > Eccesso totale di  $\text{Na}^+$

Eccesso ECF

- Scompenso cardiaco
- Cirrosi
- Sindrome nefrosica

Sodiuria  
< 10  
mmol/l

Soluzione Salina  
Ipertonica

IR  
acuta e  
cronica  
**SIADH**

Sodiuria  
> 20  
mmol/l

# IPONATREMIA

LIEVE

(135-120 mmol/l)

Asintomatico

Astenia lieve

Vertigini

MODERATA

(120-110 mmol/l)

Astenia marcata

Cefalea

Nausea

Vomito

SEVERA

(< 110 mmol/l)

Ottundimento

Allucinazioni

Epilessia

Coma

Respiro di  
Cheyne-Stokes



Iperpnea

Apnea

# TERAPIA IPONATREMIA

## Calcolo del deficit del sodio

Sapendo che l'acqua corporea totale = 60%

Uomo 70 kg con  $[Na]_p = 120$ ;  $TBW = 42L$ ;

quindi il deficit di  $Na^+ = 140 - 120 = 20 \text{ mEq} \times 42L = 840 \text{ mEq}$ .

Considerando una infusione di Salina Iperonica al 3% (513 mEq/L);  $700 \text{ mEq} / 513 = 1,6 \text{ L}$  ;

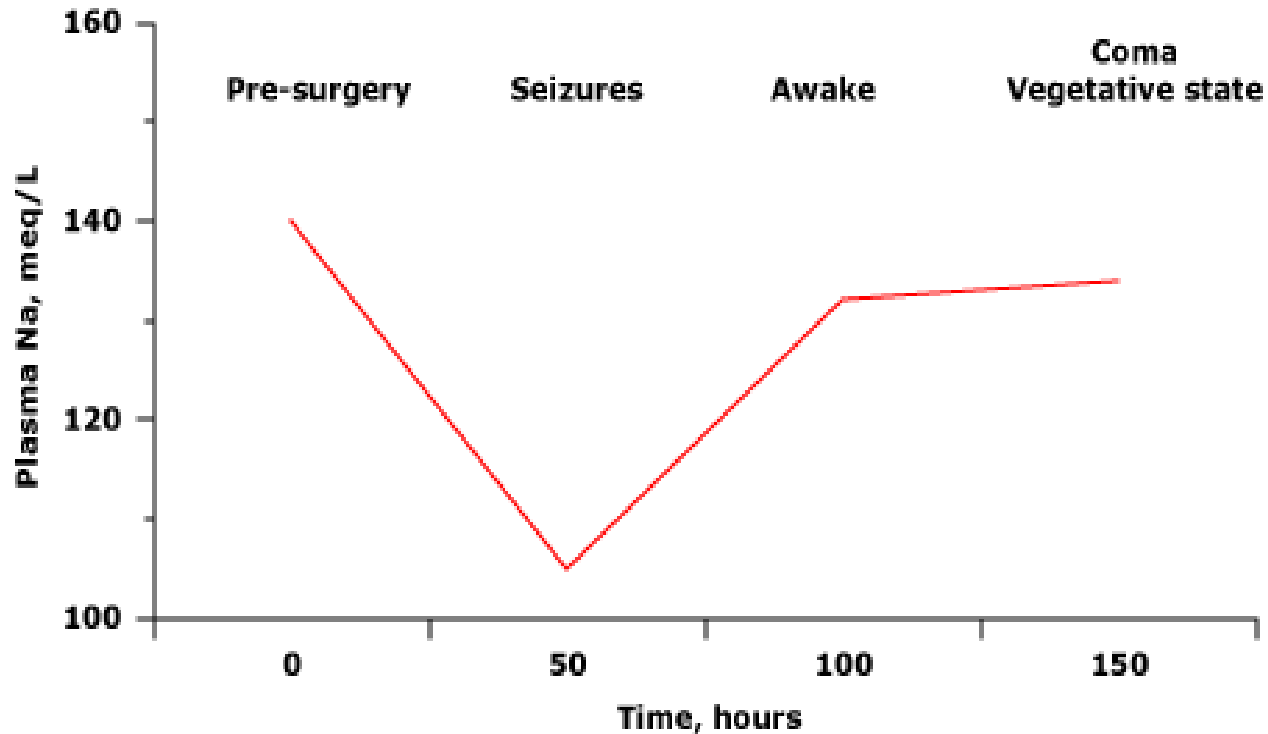
## Take Home Message:

Velocità di correzione pari a 0.5 mEq/h;

quindi 20mEq in 40h; quindi 1,6 l Iperonica in 40h ;

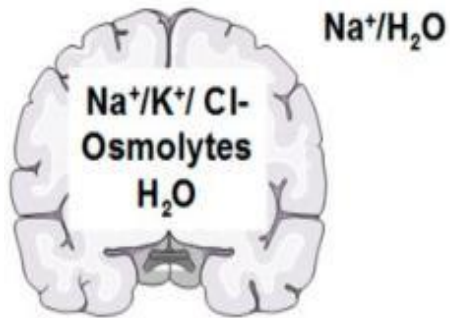
quindi =  $40\text{ml/h} / 70\text{kg} = 0,5 \text{ ml/kg/h}$

# Course of osmotic demyelination in hyponatremia

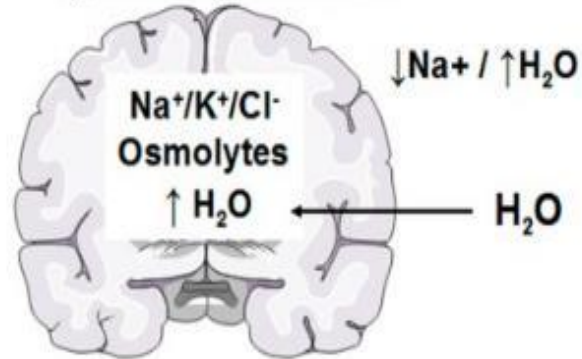


# Trattamento

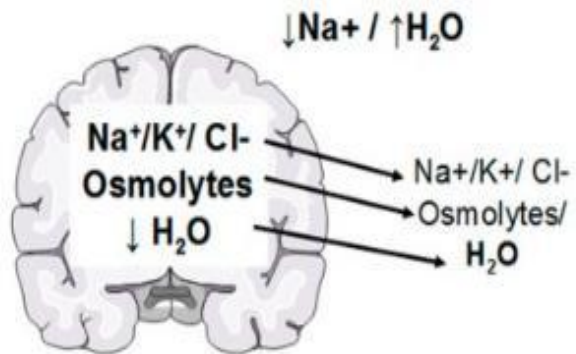
a) Normonatremia



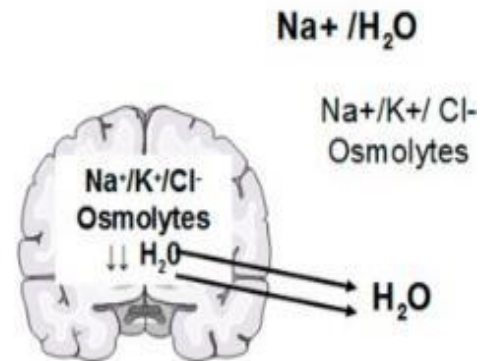
b) Acute hyponatremia



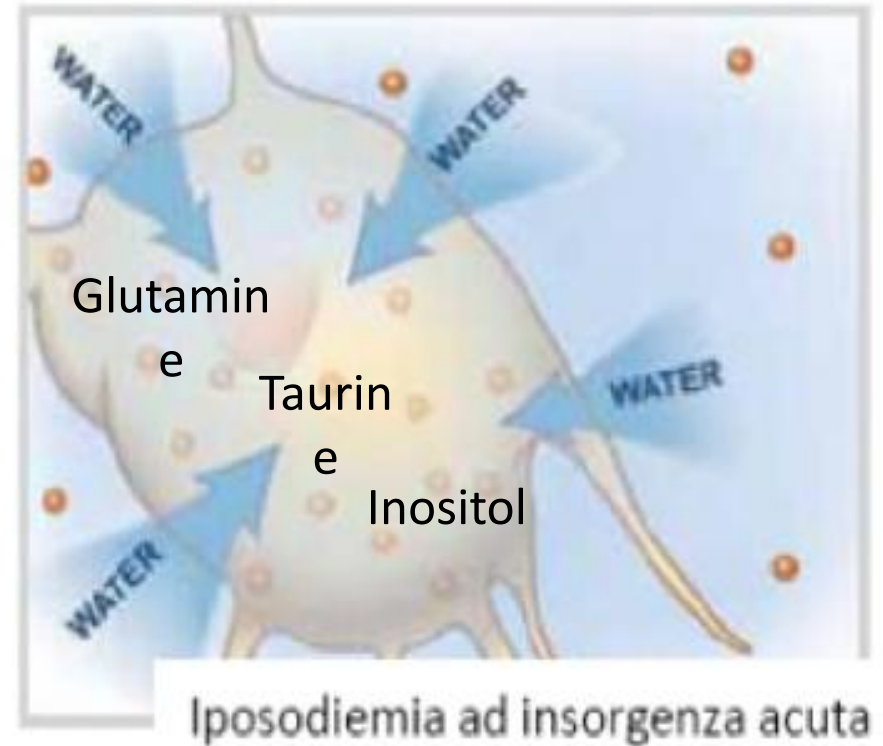
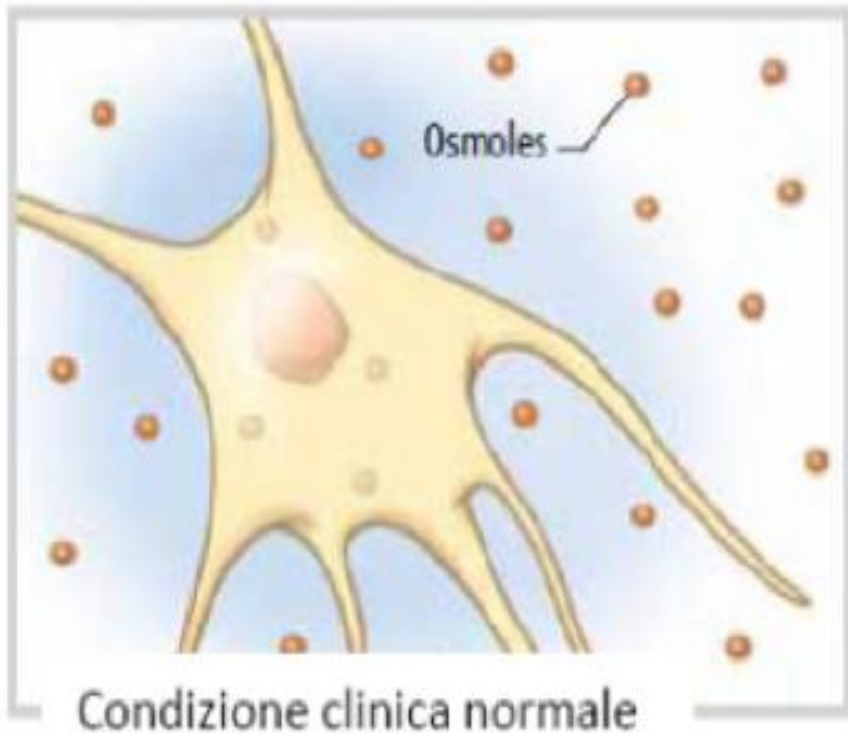
c) Chronic hyponatremia



d) Osmotic demyelination



# Stress Osmotico della Cellula Cerebrale



Iponatremia Acuta  
(swelling)

Sterns NEJM 2015

# IPONATREMIA

## Raccomandazioni



Si consiglia di correggere la sodiemia ad una **velocità < 0.5 mmol/h**

Un aumento della concentrazione sierica di sodio > 20 mmol/l in 24 h è associata ad una aumentato rischio di mielinolisi pontina (sindrome demielinizzazione osmotica)



# GRAZIE

Augiero

Barbato

Cortile

Di Nuzzo

Langella

Lo Priore

Sommese

Vernoni



Carlino – Bologna – Ciarambino - D'Addio – Guerrera - Ricciotti

Battista

Bicoku

Bottone

Campanile

D'Arco

De Vita

Di Sette

Fischetti

Gaudino

Giaquinto

Latini

Menna

Milione

Palma

Persiano

Scarano

Sena

Schettini

Volpe