

Managing Fluid Overload in Peritoneal Dialysis

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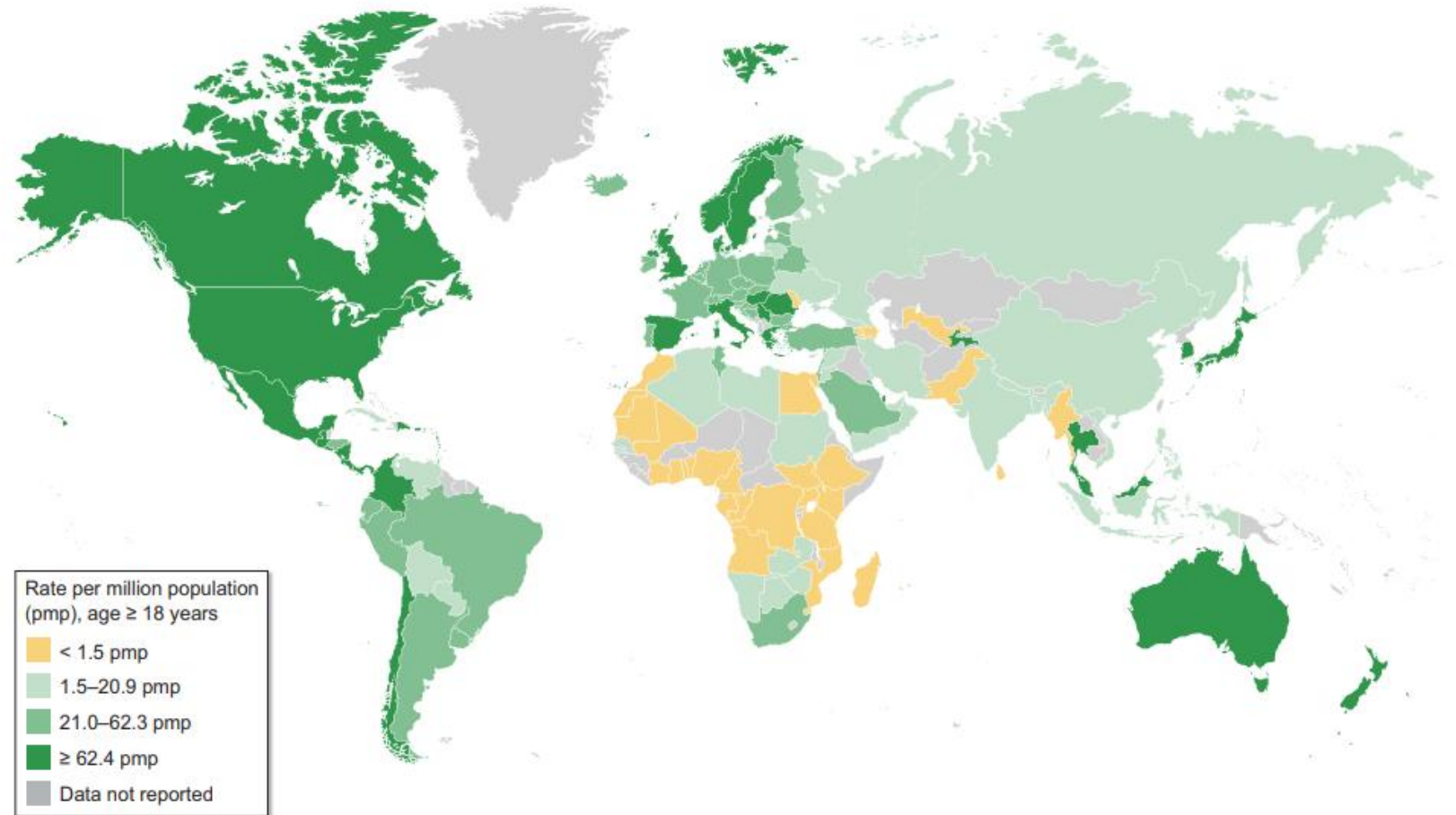
Disclaimers

- I am a nephrologist
- Participate and lead investigator-initiated trials at the Australasian Kidney Trials Network, at the University of Queensland, Australia
- Received research grants and speaker's honoraria from Vantive (Baxter) Healthcare and Fresenius Medical Care
- Current recipient of NHMRC Emerging Leadership Level 2 Investigator Grant
- Home dialysis enthusiast - especially PERITONEAL DIALYSIS

Learning Objectives

- Understand pathophysiology of fluid overload in PD
- Identify clinical consequences and risk factors
- Review assessment tools and management strategies

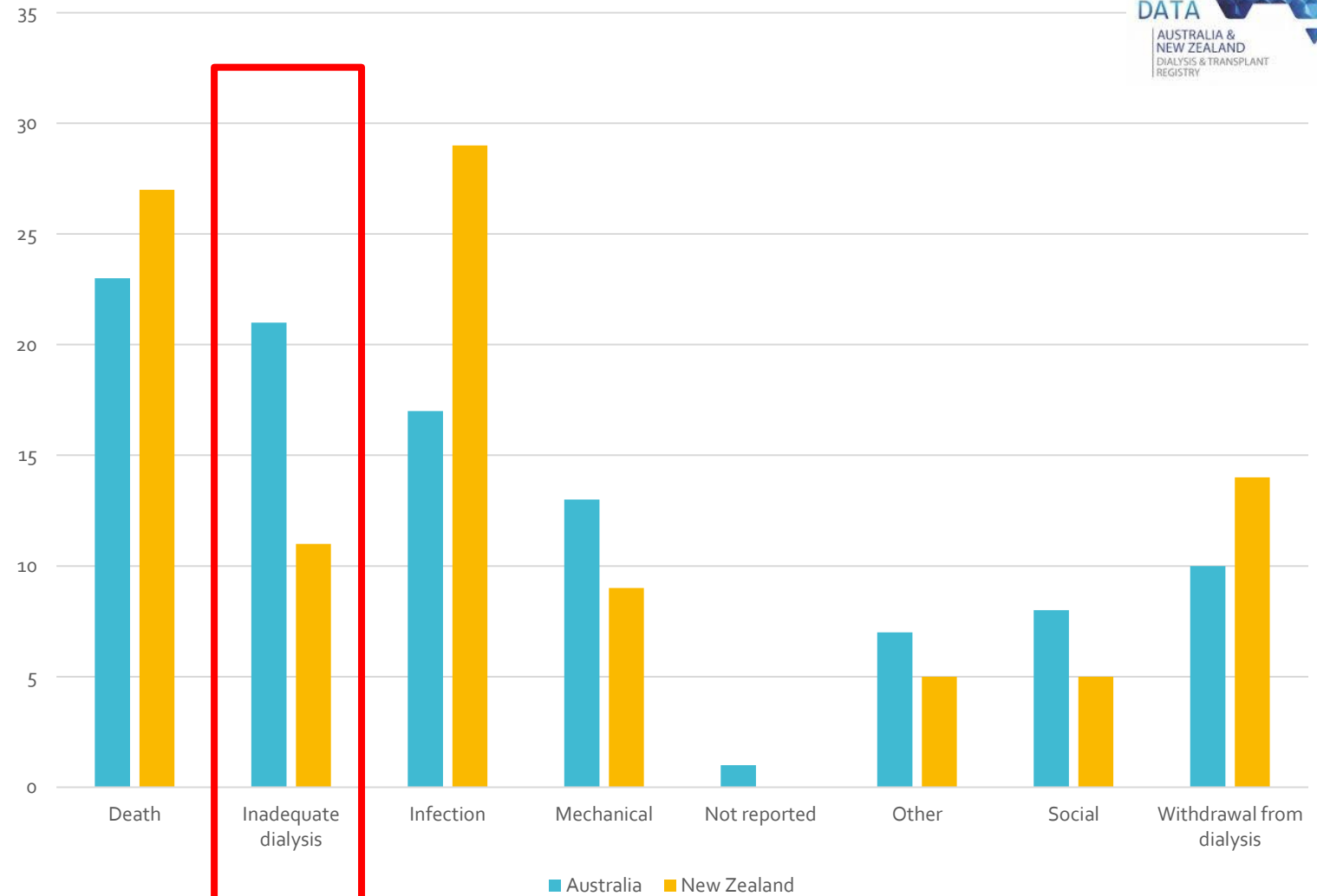
PD: Global Landscape



Median incidence of PD 22.4 pmp (prev 20.8 pmp)
Median prevalence of PD 21 pmp

**HD prevalence
322.7 pmp**

Reasons for PD cessation





“Fluid overload is common.”

And may be more common than what we think..

How reliable is clinical assessment of fluid overload?

107 patients on HD – mid-week - asymptomatic

TABLE 1 Demographic, clinical, and laboratory parameters of patients according to the tertile of BIS-derived predialysis FO

| Parameter | Overall | Low tertile (FO: <0.9 L) | Medium tertile (FO: 0.9–2.2 L) | High tertile (FO: >2.2 L) | p value |
|-----------|---------|--------------------------|--------------------------------|---------------------------|---------|
|-----------|---------|--------------------------|--------------------------------|---------------------------|---------|

TABLE 3 Univariate and multivariate linear regression analysis of risk factors associated with BIS-derived predialysis overhydration

| Risk factor | Univariate analysis | | Multivariate analysis | |
|---|----------------------|---------|-----------------------|---------|
| | Standardized β | p value | Standardized β | p value |
| Continuous variables | | | | |
| Age (years) | −0.138 | 0.15 | 0.022 | 0.81 |
| BMI (kg/m ²) | −0.243 | 0.01 | −0.158 | 0.08 |
| Interdialytic weight gain (kg) | 0.108 | 0.27 | | |
| Hemoglobin (g/dl) | −0.146 | 0.14 | −0.002 | 0.98 |
| Serum albumin (g/dl) | −0.230 | <0.05 | −0.149 | 0.11 |
| Urea reduction ratio (%) | −0.022 | 0.83 | | |
| Home SBP (mmHg) | 0.306 | 0.001 | 0.287 | <0.01 |
| Categorical variables | | | | |
| Gender (male vs. female) | 0.003 | 0.97 | | |
| Dialysis vintage (per month) | 0.358 | <0.001 | 0.306 | <0.001 |
| History of diabetes (yes vs. no) | −0.079 | 0.42 | | |
| History of CAD (yes vs. no) | −0.030 | 0.76 | | |
| History of CHF (yes vs. no) | 0.115 | 0.20 | | |
| Crackles in pulmonary auscultation (yes vs. no) | 0.081 | 0.42 | | |
| Pedal edema (yes. vs. no) | 0.094 | 0.34 | | |
| Antihypertensive drug use (yes vs. no) | 0.051 | 0.60 | | |
| Average number of antihypertensives | 0.250 | <0.01 | 0.099 | 0.29 |

Abbreviations: BMI, body mass index; CAD, coronary artery disease; CHF, congestive heart failure; SBP, systolic blood pressure.

Abbreviations: BP, blood pressure; CAD, coronary artery disease; CHF, congestive heart failure; DBP, diastolic blood pressure; OH, overhydration; SBP, systolic blood pressure.

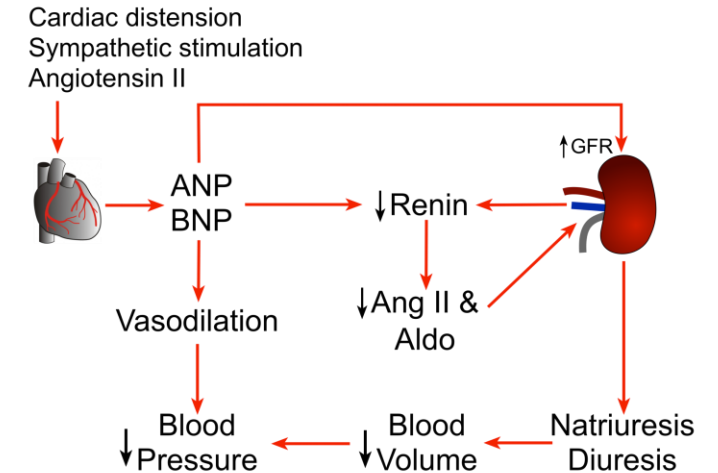
Other methods to assess fluid status

- **Serum Biomarkers**

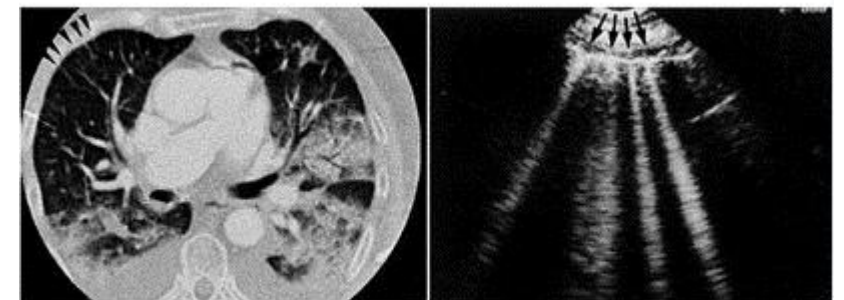
- B type natriuretic peptide (BNP)
 - Mainly reflects filling pressure of the left atrium – i.e. cardiac congestion (reflective of cardiac dysfunction and volume overload)

- **Lung Ultrasound**

- Can be used to assess the extravascular content of the lungs
- Reflects pulmonary wedge pressure – i.e., LV preload and circulating volume in relation to cardiac function rather than just fluid status



Acute pulmonary edema



Chest Computerized
Tomography

Chest Ultrasound

<https://cvphysiology.com/blood-pressure/bpo17>

Picano E et al, JASEcho, 2006; 19(3): 356-63

Bioimpedance Measurement

- Uses a flow of electrical current through the body tissues to assess body composition based on measured impedance (a compound of resistance and reactance)

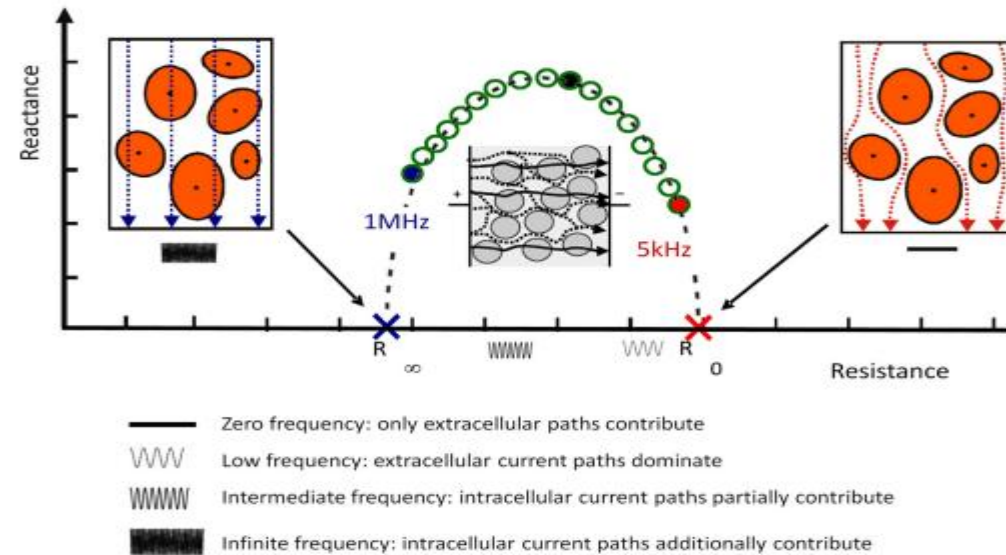


Figure 3. Principles of bioimpedance and bioimpedance spectroscopy.

- Model presume that the body is composed of 3 linked cylinders – arm, leg and the trunk (assumes constant fat-free mass – which is inaccurate in higher BMI)
- Resistance is related to water content whereas reactance is related to integrity of the cell membrane

Fluid Status in Peritoneal Dialysis Patients: The European Body Composition Monitoring (EuroBCM) Study Cohort

Wim Van Biesen^{1*}, John D. Williams², Adrian C. Covic³, Stanley Fan⁴, Kathleen Claes⁵, Monika Lichodziejewska-Niemierko⁶, Christian Verger⁷, Jurg Steiger⁸, Volker Schoder⁹, Peter Wabel⁹, Adelheid Gauly⁹, Rainer Himmele⁹, on behalf of the EuroBCM study group

- Cross sectional, observational, multi-centre study in 28 centres in 6 European countries
- To analyse hydration status in prevalent PD patients by bio-impedance spectroscopy:
 - Absolute Δ Tissue Hydration (A Δ TH) is the tissue water detected by the BCM minus the water present in the tissue under normal conditions.
- Of 639 patients included:
 - **53.4%** met 'over-hydration' criteria (A Δ TH > 90th percentile of normal population)
 - Severe fluid overload, defined as a relative change in tissue hydration above 15% was present in 25.2% of the study population (meaning ~ **1 in 4** PD patients have severe fluid overload!)

? Problem of Prevalence –

~40% of patients with <500mL/day urine output

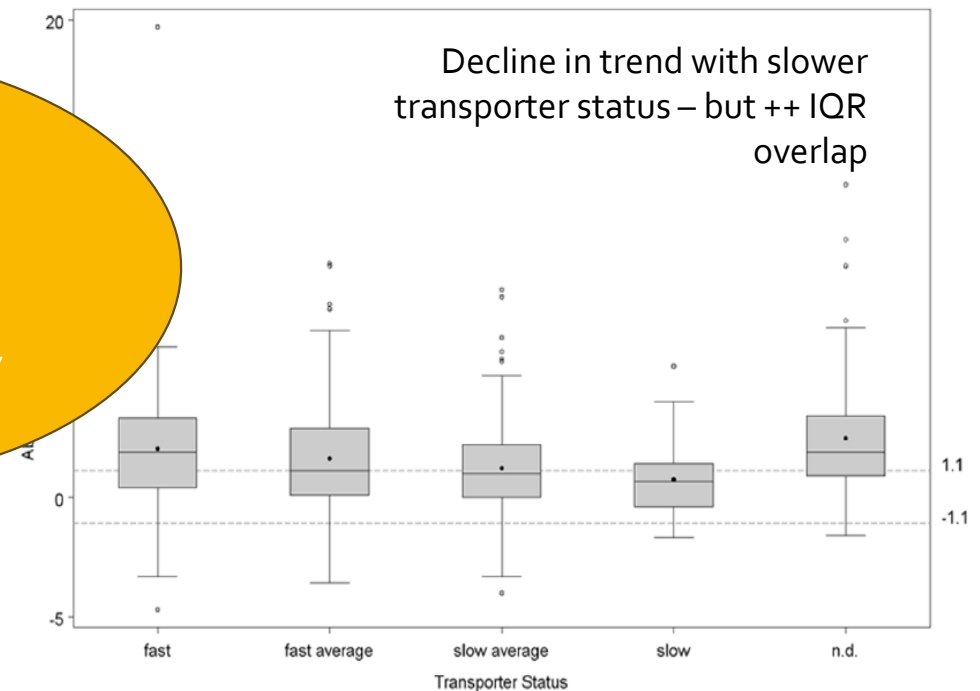


Figure 2. Box and whisker plots (median, 25th and 75th quartile, outliers) of Absolute Δ Tissue Hydration (in liters) in the different transport categories. n.d.: no peritoneal transport characteristics available in the 4 months before the BCM measurement. doi:10.1371/journal.pone.0017148.g002

Baseline hydration status in incident peritoneal dialysis patients: the initiative of patient outcomes in dialysis (IPOD-PD study)[†]

- Longitudinal, international, prospective observational study of incident PD patients
- 1092 incident patients (58.1% male) from 135 centres in 35 countries (Asia, Europe, Latin America)
- 56.4% were 'overhydrated' using BCM

Table 1. Baseline patient characteristics of the entire analysis cohort and according to hydration status

| | Total, N = 1031 | Dehydrated, N = 50 (4.8%) | Normohydrated, N = 399 (38.7%) | Overhydrated, N = 582 (56.5%) |
|------------------------------------|-----------------------|---------------------------|--------------------------------|-------------------------------|
| Age [years] | 58.0 ± 15.3 | 55.1 ± 15.1 | 56.0 ± 16.1 | 60.0 ± 14.6 |
| Gender (men) [%] | 58.1 | 42.0 | 44.6 | 68.7 |
| Height [cm] | 169.1 ± 9.1 | 164.8 ± 9.4 | 163.9 ± 9.5 | 167.8 ± 10.3 |
| Weight [kg] | 75.0 ± 14.1 | 79.8 ± 18.4 | 69.7 ± 15.5 | 73.0 ± 16.2 |
| Ethnicity [%] | | | | |
| White | 66.0 | 66.0 | 74.2 | 69.1 |
| Black | 8.0 | 8.0 | 3.3 | 3.1 |
| Asian | 12.0 | 12.0 | 12.8 | 17.9 |
| Hispanic | 6.0 | 6.0 | 4.3 | 3.6 |
| Other | 8.0 | 8.0 | 5.5 | 6.4 |
| Unknown | 0.0 | 0.0 | 17.5 | 36.9 |
| Missing | 0.0 | 0.0 | 22.1 | 16.5 |
| Missing | 0.0 | 0.0 | 17.3 | 16.2 |
| Missing | 0.0 | 0.0 | 12.3 | 7.0 |
| Missing | 16.0 | 16.0 | 11.5 | 10.0 |
| Missing | 12.0 | 12.0 | 7.5 | 6.4 |
| Missing | 2.0 | 2.0 | 7.3 | 8.6 |
| Missing | 4.0 | 4.0 | 4.5 | 7.7 |
| Missing | 2.0 | 2.0 | 2.3 | 4.3 |
| Missing | 1.5 | 1.5 | 0.8 | 1.5 |
| Missing | 6.1 | 6.1 | 6.2 | 6.9 |
| Blood pressure [mmHg] | | | | |
| Male (sys) | 142.3 ± 21.5 | 136.3 ± 23.0 | 138.1 ± 20.4 | 144.6 ± 21.6 |
| Male (dias) | 80.7 ± 13.0 | 81.8 ± 10.6 | 79.5 ± 11.9 | 81.1 ± 13.5 |
| Female (sys) | 135.6 ± 21.8 | 136.0 ± 18.5 | 131.4 ± 20.2 | 140.6 ± 23.0 |
| Female (dias) | 79.1 ± 12.5 | 80.3 ± 13.1 | 78.7 ± 12.2 | 79.5 ± 12.8 |
| Residual renal function | | | | |
| Residual urinary output [mL] | 1551 ± 753 | 1834 ± 900 | 1601 ± 752 | 1492 ± 734 |
| eGFR [mL/min/1.73 m ²] | 10.8 ± 13.3 (N = 450) | 8.7 ± 4.7 (N = 24) | 11.7 ± 15.2 (N = 171) | 10.4 ± 12.4 (N = 255) |
| Creatinine clearance [mL/min] | 10.4 ± 6.0 (N = 657) | 12.0 ± 7.1 (N = 35) | 11.8 ± 6.9 (N = 253) | 9.4 ± 4.9 (N = 369) |
| Urea clearance [mL/min] | 5.4 ± 3.7 (N = 453) | 6.1 ± 4.9 (N = 24) | 5.7 ± 3.5 (N = 173) | 5.2 ± 3.7 (N = 256) |
| Extracellular water [L] | 17.3 ± 4.0 | 16.4 ± 4.2 | 15.3 ± 3.1 | 18.8 ± 3.9 |
| Intracellular water [L] | 18.6 ± 4.7 | 22.6 ± 7.1 | 18.0 ± 4.4 | 18.7 ± 4.5 |
| ECV/ICV | 0.9 ± 0.2 | 0.7 ± 0.1 | 0.9 ± 0.1 | 1.0 ± 0.2 |
| FO [L] | 1.9 ± 2.4 | -1.8 ± 0.6 | 0.2 ± 0.6 | 3.3 ± 2.08 |

When in doubt – fluid overload is more likely than not!

Target Weight in PD = normotensive, euvolaemic and symptom free

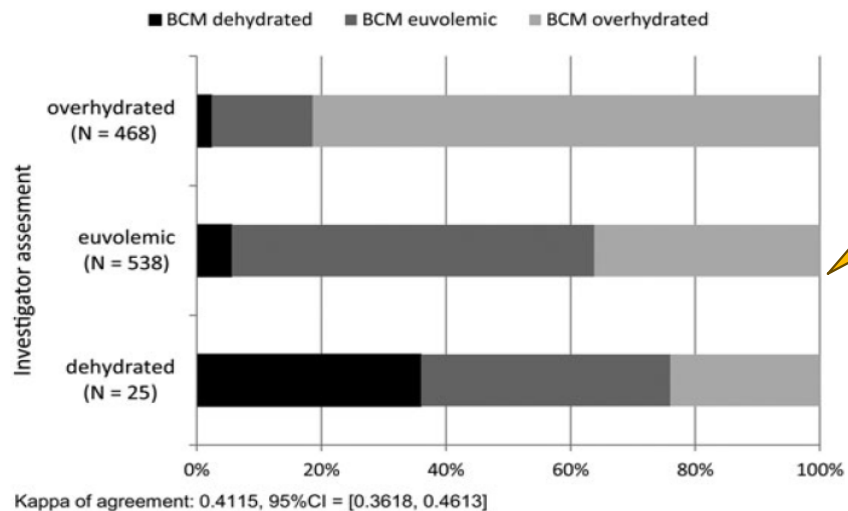


FIGURE 4: Assessment of hydration status with BCM versus clinical assessment by investigator using the physician specific assessment.

“Fluid overload is common and associated with adverse patient outcomes”

SONG-PD



1 CORE OUTCOMES

Critically important to all stakeholder groups
Report in all trials

2 MIDDLE TIER




Critically important to some stakeholder groups
Report in some trials

3 OUTER TIER

Important to some or all stakeholder groups
Consider for trials



The impact of volume overload on technique failure in incident peritoneal dialysis patients

François Vrtovsnik¹, Christian Verger ², Wim Van Biesen ³, Stanley Fan⁴, Sug-Kyun Shin⁵, Carmen Rodríguez⁶, Isabel Garcia Méndez⁷, Frank M. van der Sande⁸, Tatiana De los Ríos⁹, Katharina Ihle⁹, Adelheid Gauly ⁹, Claudio Ronco¹⁰ and James Heaf¹¹, for the IPOD-PD Study Group

- From IPOD-PD Study, 719 patients meeting inclusion criteria were included (e.g., f/up >6 months, valid measurement of BCM data at month 6).
- BCM was measured at baseline and every 3 monthly.
- Outcome: time to technique failure (defined as composite of death + t/f to HD)
- Transplantation was considered a competing event

Table 1. Patient characteristics at baseline (technique failure analysis population, n = 719)

| Characteristics | Technique failure analysis population (n = 719) |
|--|---|
| Age, years | 58.4 ± 14.9 |
| Sex (male), % | 57.2 |
| Height, cm | 166.9 ± 10.1 |
| Weight, kg | 73.3 ± 16.3 |
| Blood pressure (systolic), mmHg | 137.9 ± 22.7 |
| Blood pressure (diastolic), mmHg | 80.2 ± 12.6 |
| Transport status (first assessment within 6 months), % | |
| High (fast) | 8.8 |
| High average | 28.5 |
| Low average | 20.2 |
| Low (slow) | 16.7 |
| Missing | 25.9 |
| Primary renal disease, % | |
| Diabetes | 21.1 |
| Glomerulonephritis | 21.1 |
| Hypertension | 12.8 |
| Hereditary/congenital disease | 12.2 |
| Other | 18.9 |
| Unknown | 13.8 |
| Comorbidities, % | |
| Hypertension | 89.0 |
| Diabetes (Types 1 + 2) | 37.4 |
| Liver disease | 4.8 |
| Cardiovascular disease (NYHA Stages I, II, III, IV, unknown) | 24.1 (7.9, 6.7, 3.1, 1.0, 5.4) |

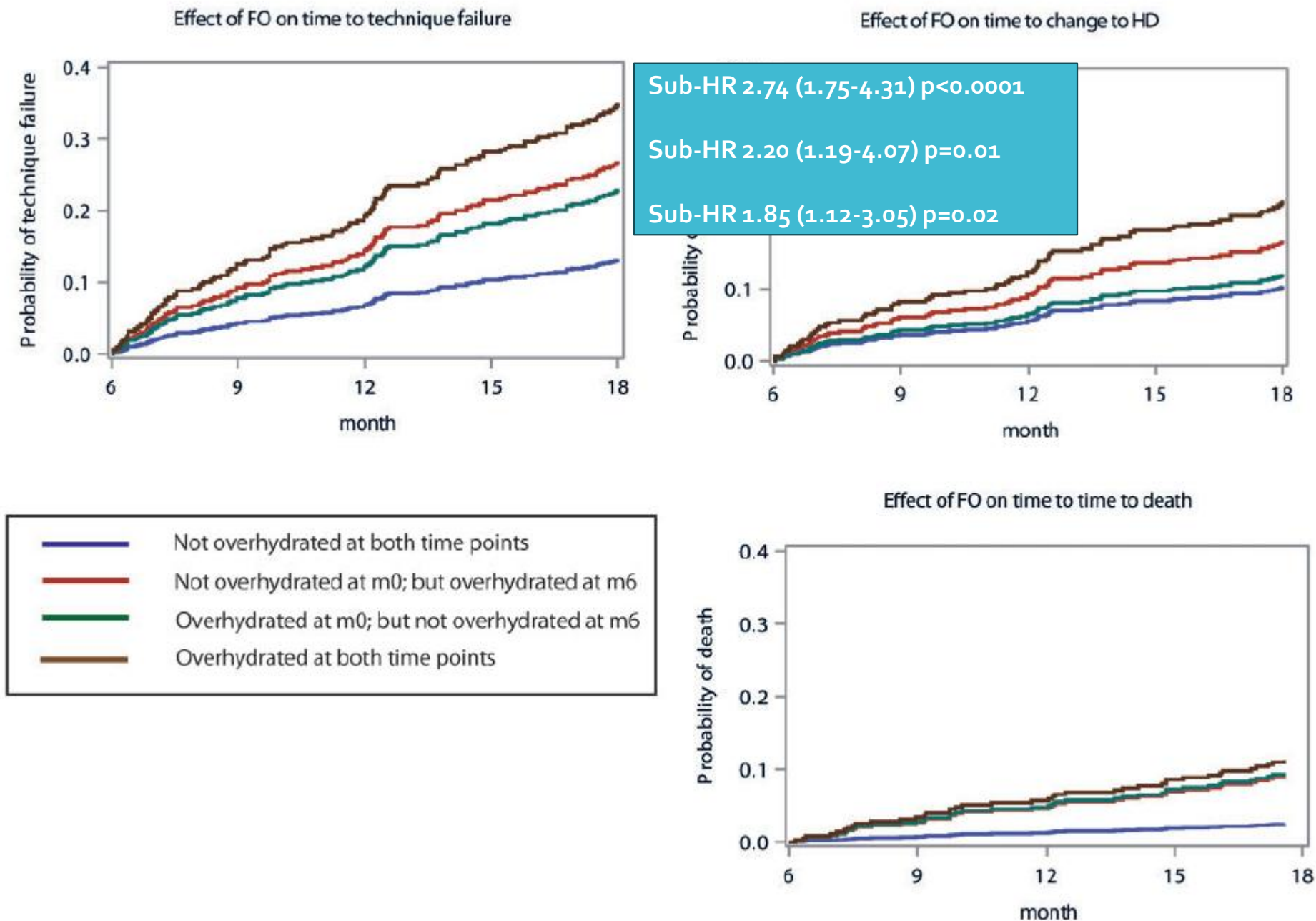


FIGURE 2: Cumulative incidence curves displaying the effect of fluid overload on composite endpoint and the two single components transfer to HD and death. m0, Month 0; m6, Month 6; FO, fluid overload.

Asymptomatic fluid overload predicts survival and cardiovascular event in incident Chinese peritoneal dialysis patients

Jack Kit-Chung Ng, Bonnie Ching-Ha Kwan, Kai-Ming Chow, Wing-Fai Pang, Phyllis Mei-Shan Cheng, Chi-Bon Leung, Philip Kam-To Li, Cheuk-Chun Szeto*

- Single-centre study of 311 incident PD patients, median follow-up of 27.3 months.
- Volume status measured by BIS:
 - Volume of Overhydration (OH)
 - OH/extracellular water (ECW) ratio
 - ECW/total body water (TBW) ratio
 - ECW to intracellular water (ICW) ratio (E:I ratio)
- Outcomes:
 - Patient survival
 - Technique survival
 - cardiovascular event-free survival

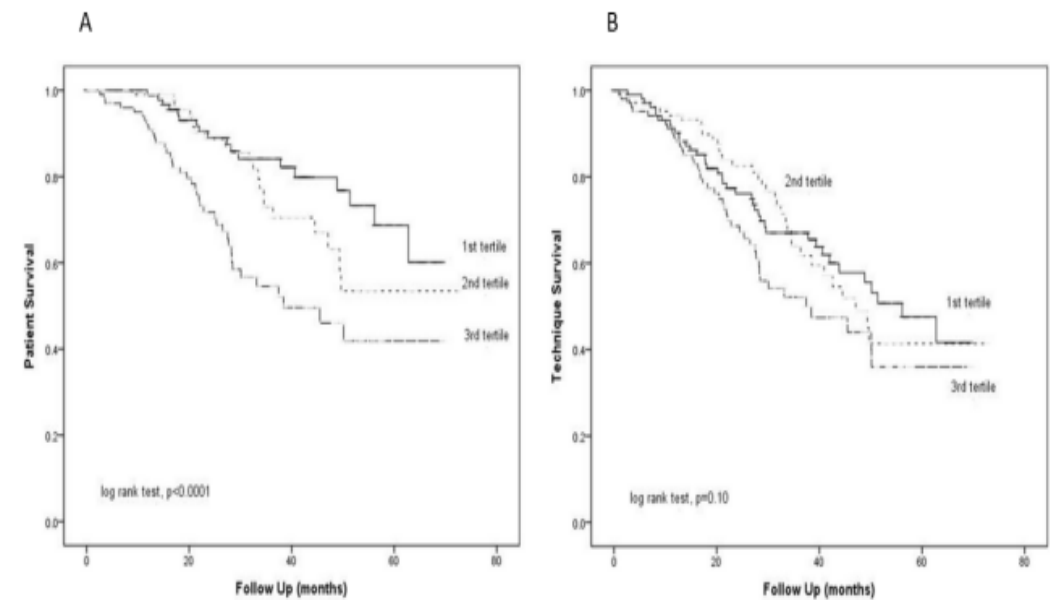


Fig 1. Kaplan-Meier plot of (A) patient survival; and (B) technique survival. Patients were divided into tertiles according to their baseline E:I ratio (1st tertile: ≤ 0.91 ; 2nd tertile > 0.91 – 1.07 ; 3rd tertile > 1.07). Data were compared by the log rank test.

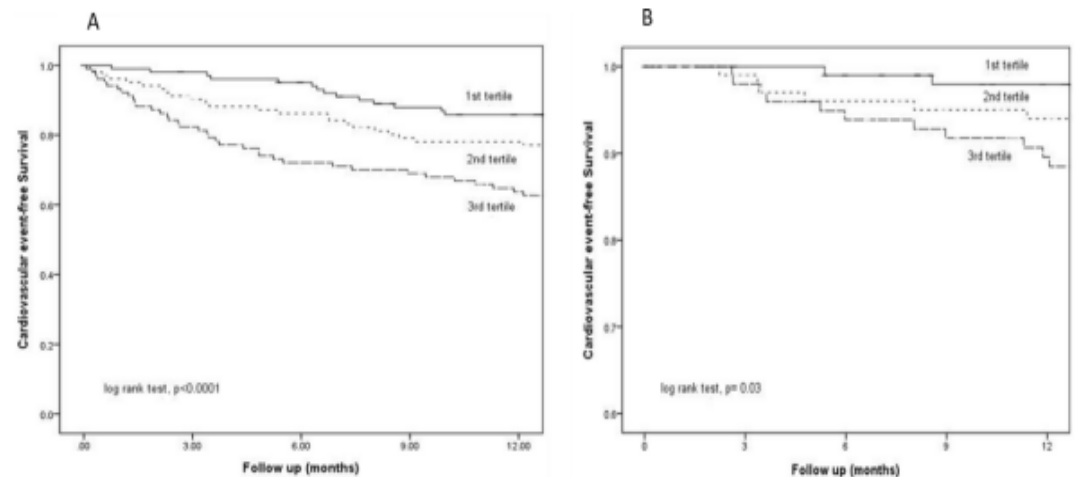


Fig 2. Kaplan-Meier plot of cardiovascular event-free survival with hospital admission for congestive heart failure (A) included; and (B) excluded. Patients were divided into tertiles according to their baseline E:I ratio (1st tertile: ≤ 0.91 ; 2nd tertile > 0.91 – 1.07 ; 3rd tertile > 1.07). Data were compared by the log rank test.



KEEP
CALM
AND
DON'T
PANIC

~~PROBLEM~~
SOLUTION



Understand
the CAUSE/s
to inform
strategies of
ACTION



Inadequate peritoneal
ultrafiltration

Reduced residual
kidney function

Increased dietary salt
and fluid intake

Fluid Restriction

- Generally, “restrictive” (e.g., 1.5L/day)
- NHS Oxford Kidney Unit patient information brochure for PD states:
 - Your fluid allowance = 750mL + previous days 24-hour urine output

How much fluid you can have differs from person to person. This depends on kidney function, urine output, and dialysis. Your fluid allowance may also change over time. Discuss with your kidney doctor, nurse, or dietitian if you are unsure of your allowance.



My fluid allowance is _____ mL per day.

Let's talk about fluid – 'obvious' and 'hidden'

| Example 1 | | Example 2 | |
|---|---------------|--|---------------|
| Breakfast | Fluid mL | Breakfast | Fluid mL |
| 2 slices of wholegrain bread with 2 poached eggs / tomato / mushroom / jam / honey | | ½-1 cup high fibre cereal with 1/2 cup low fat milk | 125 |
| 1-2 tablespoons dried fruit | | 1 medium sized orange | 80 |
| 1 cup tea / coffee / water | 250 | ½ cup tea / coffee / water | 125 |
| Breakfast total = 250mL | | Breakfast total = 330mL | |
| Lunch | | Lunch | |
| 2 slices wholegrain bread or 1 x wrap or 4 rice cakes with scrape of avocado / marg / butter | | 1 cup salt reduced soup +/- bread | 250 |
| with cheese, 1 cup mixed salad +/- lean meat / egg / tuna / salmon / chicken | | 1 tub of low fat / diet yoghurt (150g) | 120 |
| ½ cup tinned fruit | 125 | ½ cup tea / coffee / water / juice | 125 |
| 1 cup tea / coffee / water / juice | 250 | | |
| Lunch total = 375mL | | Lunch total = 495mL | |
| Dinner | | Dinner | |
| 100-120g of lean meat / chicken (no skin) / fish | | 100-120g of lean meat / chicken (no skin) / fish | |
| baked potato | | with 1 cup rice / pasta (cooked) | 100 |
| plenty of other mixed cooked vegetables / salad | | plenty of other mixed cooked vegetables / salad | |
| low fat/diet yoghurt (150g) | 120 | ½ cup custard | 100 |
| 1 cup tea / coffee / water | 250 | ½ cup tea / coffee / water | 125 |
| Dinner fluid total = 370mL | | Dinner fluid total = 325mL | |
| Snack Ideas | | Snack Ideas | |
| 1 cup of low fat milk | 250 | 1 piece of fruit | 80 |
| 1 thin slice of fruit bread / raw unsalted nuts (2 tablespoons) OR 2-3 grainy crackers with cottage cheese and tomato | | 1 thin slice of fruit bread / raw, unsalted nuts (2 tablespoons) OR 2-3 grainy crackers with cottage cheese and tomato | |
| Snack total = 250mL | | Snack total = 80mL | |
| Plus 1 additional cup of fluid (water / juice / soda water / tea / coffee) | 250 | Plus 1 additional cup of fluid (water / juice / soda water / tea / coffee) | 250 |
| TOTAL Example 1 | 1495mL | TOTAL Example 2 | 1480mL |

Control your THIRST

SALT RESTRICTION

- <2g (87mmol/L) / day recommended in PD
- Strategies:
 - Fresh, minimally processed foods
 - Remove salt on table
 - Cook with little/no added salt
 - Use herbs/spices to flavour
 - Limit the use of commercial sauces, dressings and instant products
 - Choose foods with lower sodium content (e.g., <120mg/100g of sodium = best)

GLYCAEMIC CONTROL

- Diet
- PD Prescription – lower glucose strength
- Glucose-lower treatments
 - OHG – DPP₄ inhibitors (e.g., linagliptin)
 - Insulin – tailored to individual needs (consider PD prescription); multiple daily injection preferred (mix of long- and short-actings) – if unable premixed 70/30 or 70/25

| Food Groups | OK to Eat <i>Choose these foods every day. One serving has less than 100 mg sodium.</i> | Eat Sometimes <i>Choose these foods once in a while. One serving has less than 300 mg sodium.</i> | Avoid These Foods <i>Choose these foods rarely. One serving has more than 300 mg of sodium.</i> |
|---|---|--|---|
| Fruits | <ul style="list-style-type: none"> • All fruits and fruit juices | | |
| Vegetables | <ul style="list-style-type: none"> • All fresh, frozen, and canned vegetables with no salt or sauces added | <ul style="list-style-type: none"> • Most canned vegetables with salt • Low-sodium tomato and V-8 juice | <ul style="list-style-type: none"> • Regular tomato or V-8 juice • Ready-to-eat tomato sauces • Pickled foods like sauerkraut, olives, and pickles |
| Meats and Other Protein Foods | <ul style="list-style-type: none"> • Fresh meats, poultry, and fish (no salt added) • Clams and mussels, steamed • Unsalted peanut butter and nuts • Tuna fish, canned without added salt • Dried beans or peas, cooked without salt • Tofu • Eggs | <ul style="list-style-type: none"> • Peanut butter • Shellfish, not treated with salt • Canned beans • Frozen meals with less than 300 mg sodium | <ul style="list-style-type: none"> • Tuna fish, canned with added salt • Fried foods • Salted, smoked, cured, or canned meats • Lunch deli meats • Spam • Corned beef • Hot dogs and sausage • Jerky • Ham and bacon • Frozen meals with more than 600 mg sodium |
| Dairy | <ul style="list-style-type: none"> • Unsalted butter • Unsalted cottage cheese • Milk and yogurt | <ul style="list-style-type: none"> • Buttermilk • Lower sodium cheeses like mozzarella, Swiss, cheddar, ricotta | <ul style="list-style-type: none"> • High-sodium cheeses like feta, bleu, and goat cheese • Cottage cheese |
| Cereals, Grains, and Starches | <ul style="list-style-type: none"> • Plain hot cereal • Shredded wheat or puffed rice cereal • Unsalted corn tortillas, granola, quick breads • Low-sodium breads • Rice, pasta, whole grains, cooked without salt • Unsalted potatoes and squash | <ul style="list-style-type: none"> • Salted crackers and pretzels • Most cold cereals • Most breads and muffins • Pancakes and waffles | <ul style="list-style-type: none"> • Stuffing mixes • Cornbread • Rice or noodles prepared with flavor packets |
| Spices, Seasonings, and Condiments | <ul style="list-style-type: none"> • Fresh or dried herbs • Powders, not salts (for example, use onion and garlic powder instead of onion and garlic salt) • Tabasco (no more than 1 tablespoon) • Vinegar, lemon and lime juice • Fresh garlic • Sesame seeds • Allspice • Mrs. Dash and Lawry's salt-free seasonings • Flavored extracts like vanilla and almond | <ul style="list-style-type: none"> • Some hot sauces (check label for sodium content) • Mayonnaise • Mustard • Low-sodium ketchup | <ul style="list-style-type: none"> • Most canned soups • Bouillon cubes and prepared broths • Sauces: teriyaki, soy, fish, Worcestershire • Sea salt, regular salt • Baking soda (regular) • Garlic, onion, and other flavored or seasoned salts • Relishes • Monosodium glutamate (MSG) • Meat tenderizers • Ketchup and barbecue sauce • All seasoning packets • Salad dressing • Cheese sauce mixes, such as Alfredo sauce • Olives • Some hot sauces |

Work your KIDNEYS

- Diuretics should be utilised (and 1st therapeutic choice in those with preserved RKF)
- Options include:
 - Loop diuretics – frusemide (up to 500mg/day)
 - Others – HCT, metolazone

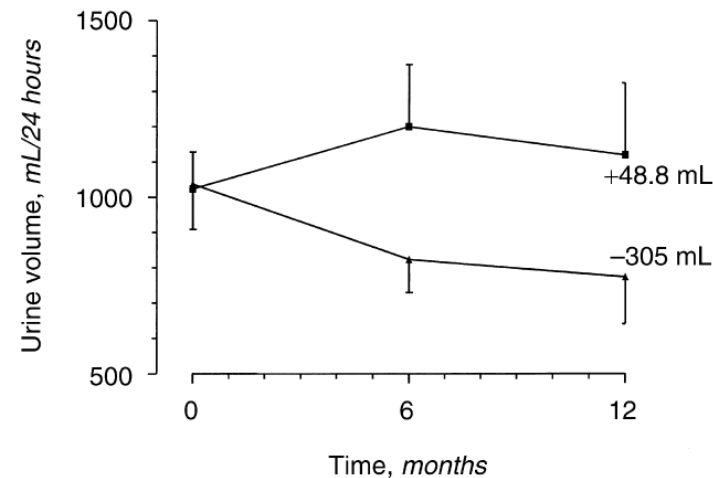


Fig. 1. Evolution of urine volume (UV) over one year of peritoneal dialysis (PD). UV at randomization was comparable between groups. In the diuretic group (■), it remained constant over one year of CAPD, whereas in the control group (▲), UV declined. Data presented are mean \pm SEM at each time point.

- 61 incident patients on CAPD randomised 1:1 to 250mg daily frusemide vs. placebo

Table 2. Change in urine volume, urinary sodium, urea and creatinine clearance

| | Control | Diuretic | <i>P</i> value |
|---|-------------------|-------------------|----------------|
| Δ Urine volume <i>mL/month</i> | -23.3 ± 11.2 | $+6.47 \pm 9.52$ | 0.047 |
| Δ 24-hour urinary sodium <i>mmol/24 h</i> | -2.57 ± 1.51 | $+0.72 \pm 0.85$ | 0.041 |
| Δ Creatinine clearance <i>mL/min/month</i> | -0.071 ± 0.04 | -0.12 ± 0.05 | 0.45 |
| Δ Urinary Kt/v <i>per month</i> | -0.019 ± 0.01 | -0.020 ± 0.01 | 0.92 |

Efficacy of triple diuretic treatment in continuous ambulatory peritoneal dialysis patients: A randomized controlled trial

Raweewan Witoon, Somchai Yongsiri, Prapan Buranaburidej, Pacharin Nanna

Table 2. Overhydration measured by BIS in the single diuretic group and the triple diuretic group at baseline, 3rd month, and 6th month of study

| | Single diuretic | Triple diuretics | <i>P</i> value |
|---|-----------------|------------------|----------------|
| OH (L) | | | |
| Baseline | 2.27 ± 2.35 | 2.94 ± 2.08 | 0.34 |
| 3rd month | 2.03 ± 1.80 | 1.03 ± 0.68 | 0.01 |
| 6th month | 2.78 ± 2.42 | 1.39 ± 1.64 | 0.06 |
| ΔOH (L) | | | |
| 3rd month vs. baseline (OH 3rd month–OH baseline) | 1.84 ± 2.27 | 0.44 ± 1.62 | 0.03 |
| 6th month vs. baseline (OH 6th month–OH baseline) | 1.49 ± 2.82 | –0.48 ± 2.61 | 0.02 |

Data are presented as mean ± standard deviation.

BIS, bioimpedance spectroscopy; OH, overhydration.

No difference in adverse events

Table 1. Baseline demographic features of both groups

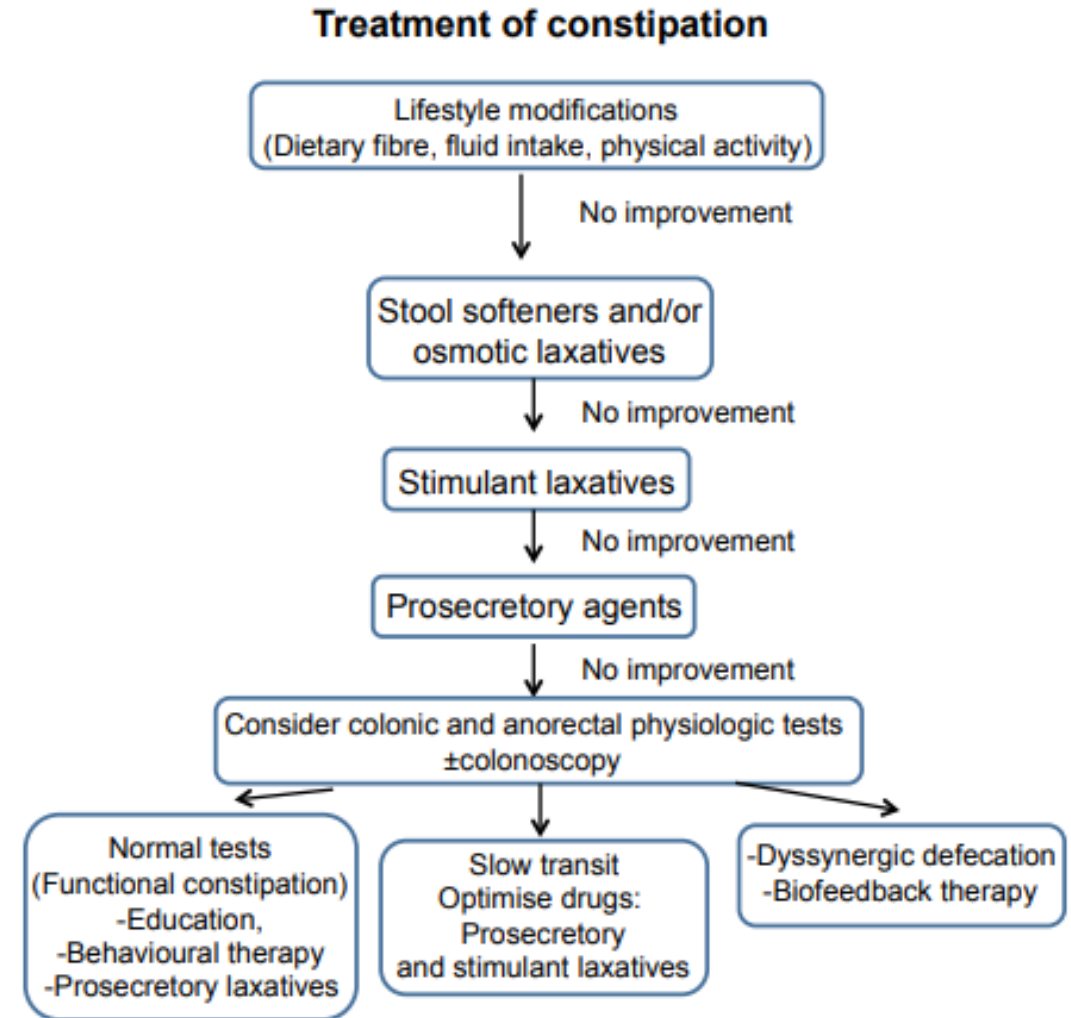
| Baseline characteristic | All patients (n = 51) | Single diuretic (n = 27) | Triple diuretics (n = 24) | <i>P</i> value |
|--|-----------------------|--------------------------|---------------------------|----------------|
| Sex, female/male | 30/21 | 18/9 | 12/12 | 0.265 |
| Age (yr) | 59.29 ± 9.81 | 59.25 ± 10.37 | 58.5 ± 6.79 | 0.768 |
| Comorbidity | | | | |
| Diabetes | 34 (66.7) | 17 (63.0) | 17 (70.8) | 0.767 |
| Hypertension | 36 (70.6) | 17 (63.0) | 19 (79.2) | 0.235 |
| Dyslipidemia | 4 (7.8) | 2 (7.4) | 2 (8.3) | 0.999 |
| Dialysis vintage (mo) | 12.19 ± 13.95 | 11.74 ± 14.42 | 17.17 ± 17.18 | 0.231 |
| Residual urine volume (mL) | 855.00 ± 508.92 | 870.74 ± 474.38 | 837.50 ± 620.53 | 0.832 |
| < 100 mL | 2 (3.9) | 0 (0) | 2 (8.3) | 0.402 |
| 100–500 mL | 19 (37.3) | 10 (37.0) | 9 (37.50) | |
| 501–1,000 mL | 9 (17.7) | 6 (22.2) | 3 (12.50) | |
| > 1,000 mL | 21 (41.2) | 11 (40.7) | 10 (41.7) | |
| Peritoneal membrane type | | | | |
| Low | 1 (2.0) | 0 (0) | 1 (4.2) | 0.061 |
| Low average | 19 (37.3) | 6 (22.2) | 13 (54.2) | |
| High average | 16 (31.4) | 11 (40.7) | 5 (20.8) | |
| High | 3 (5.9) | 1 (3.7) | 2 (8.3) | |
| No data | 12 (23.5) | 9 (33.3) | 3 (12.5) | |
| Dialysis adequacy | | | | |
| Kt/V urine | 0.64 ± 0.98 | 0.68 ± 1.22 | 0.60 ± 0.69 | 0.772 |
| Kt/V PD | 1.85 ± 0.40 | 1.91 ± 0.41 | 1.80 ± 0.38 | 0.325 |
| Kt/V total | 2.48 ± 0.99 | 2.59 ± 1.22 | 2.38 ± 0.72 | 0.453 |
| Renal CCr (L/wk/1.73 m ²) | 24.50 ± 31.42 | 20.33 ± 34.32 | 28.49 ± 28.58 | 0.359 |
| PD CCr (L/wk/1.73 m ²) | 43.57 ± 20.01 | 39.50 ± 27.10 | 47.46 ± 8.19 | 0.156 |
| Daily net glucose exposure | 139.54 ± 39.21 | 138.54 ± 25.21 | 144.21 ± 43.27 | 0.590 |
| Antihypertensive drugs | | | | |
| Angiotensin converting enzyme inhibitors | 5 (9.8) | 2 (7.4) | 3 (12.5) | 0.656 |
| Angiotensin receptor blockers | 30 (58.8) | 15 (55.6) | 15 (62.5) | 0.777 |
| Calcium channel blockers | 14 (27.5) | 10 (37.0) | 4 (16.7) | 0.127 |
| Beta blockers | 31 (60.8) | 19 (70.4) | 12 (50.0) | 0.161 |
| Others | 16 (31.4) | 10 (37.0) | 6 (25.0) | 0.385 |
| Diuretic drugs | | | | |
| Loop diuretics | 51 (100) | 27 (100) | 24 (100) | 0.999 |
| Thiazide | 0 (0) | 0 (0) | 0 (0) | 0.999 |
| Spironolactone | 1 (2.0) | 1 (3.7) | 0 (0) | 0.999 |

Make sure PD is working (optimally)

- Rule out constipation
- Ensure proper adherence to prescription
- Rule out catheter flow dysfunction
- Rule out hernias/leaks
- Optimise PD prescription with cautious increase in concentration of dextrose-based solutions as needed

Avoid Constipation

| Laxative class | Site of action | Mechanisms of action | Medication | Adverse events |
|---------------------------|-----------------------|---|---|---|
| Bulk-producing agents | Small and large bowel | Retain water in stool improving consistency and peristalsis | Psyllium Methylcellulose Dietary fiber Calcium polycarbophil | Flatulence, bloating, abdominal distension |
| Emollient stool softeners | Colon | Soften stool by promoting luminal water binding/detergent-like action | Docusate sodium and calcium | Diarrhea, Intestinal cramping |
| Stimulants | Colon | Stimulate colonic contractions/decrease intestinal water absorption | Senna Aloe Bisacodyl Sodium picosulfate Magnesium hydroxide | Abdominal discomfort and cramps, nausea, vomiting, diarrhea, rectal irritation DO NOT use if concern for obstruction |
| Osmotic laxatives | Colon | Osmotic water binding | Polyethylene glycol Lactulose Milk of magnesia Magnesium citrate/sulfate | Bloating, cramps, flatulence, electrolyte disturbance |



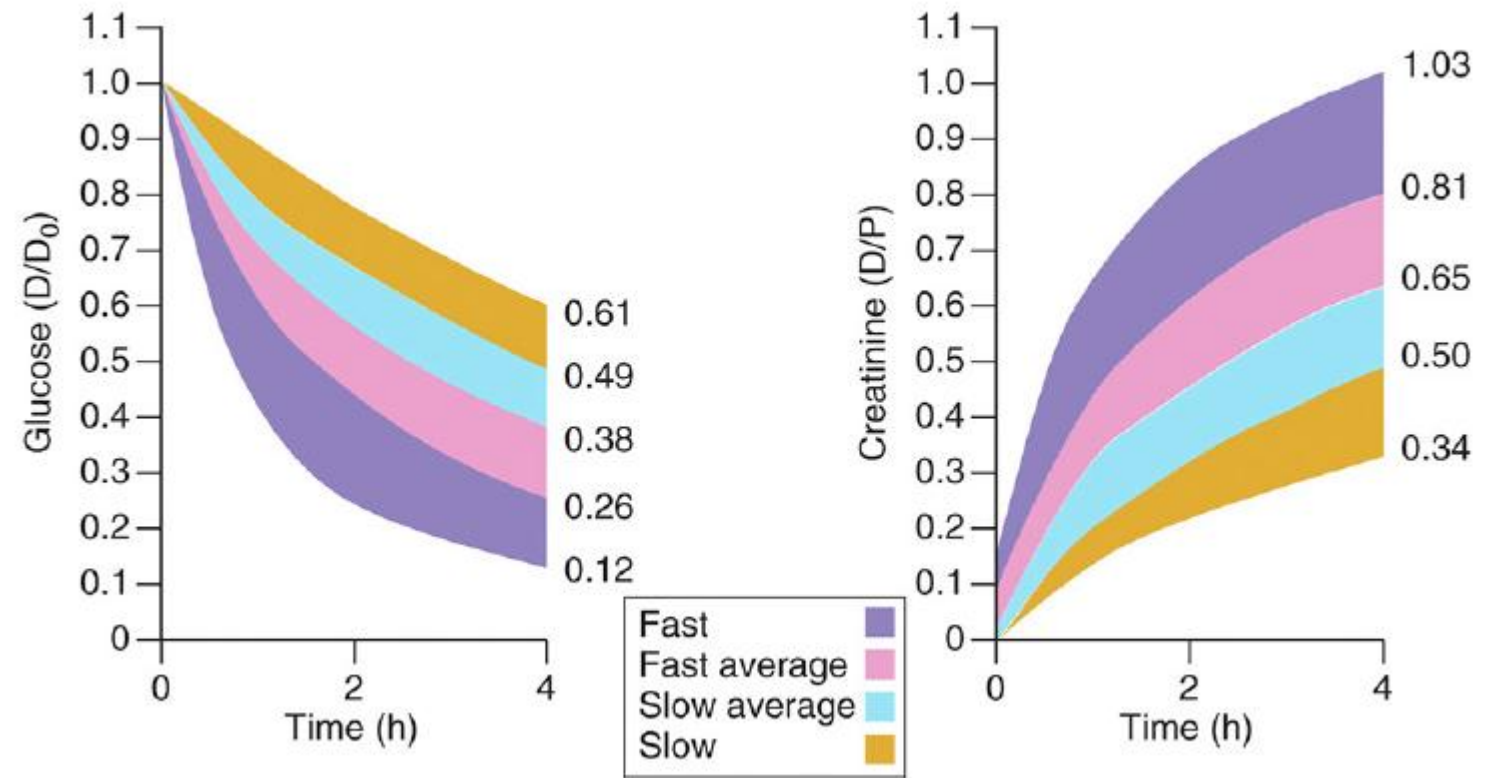
Check
Adherence to
PD and
implement
timely
intervention

Table 3:

Number of adverse events and the IRR for events in group APD versus events in group RM-APD,

| Type of event | Crude APD/RM-APD | IRR (95% CI) | $P > z $ |
|--|---------------------------------------|-------------------------|-------------|
| PD infection events | 100/86 ($P = .12$) | 1.28 (0.69–2.37) | .42 |
| Mechanical events | 33/62 ($P = .001$) | 0.42 (0.18–0.98) | .04 |
| Intra-abdominal pressure events | 6/13 ($P = .07$) | 0.48 (0.16–1.40) | .18 |
| Fluid balance events | 123/35 ($P = .001$) | 3.44 (1.95–6.07) | .001 |
| Cardiovascular events | 79/44 ($P = .02$) | 1.81 (1.03–3.09) | .03 |
| Metabolic events | 108/44 ($P = .001$) | 2.78 (1.51–5.38) | .005 |
| Other events | 53/77 ($P = .005$) | 1.01 (0.45–2.29) | .97 |
| Mechanical hospitalization events | 18/45 ($P = .004$) | 0.41 (0.22–0.66) | .001 |
| FOL and IDE hospitalizations | 19/7 ($P = .01$) | 3.24 (1.24–7.53) | .03 |
| All-cause deaths | 55/33 ($P = .01$) | 1.82 (1.18–2.81) | .006 |
| Cardiovascular deaths | 24/13 ($P = .05$) | 2.03 (1.03–3.97) | .04 |

Optimise PD Prescription



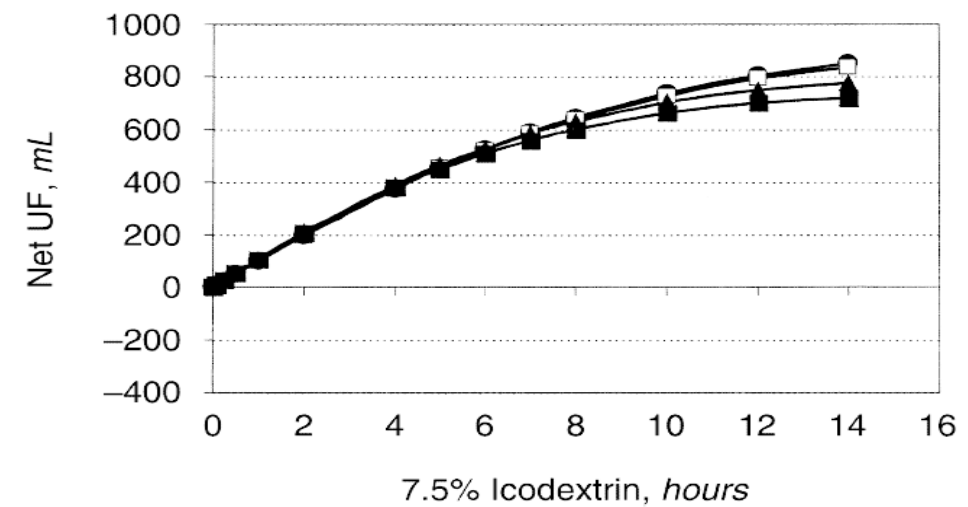
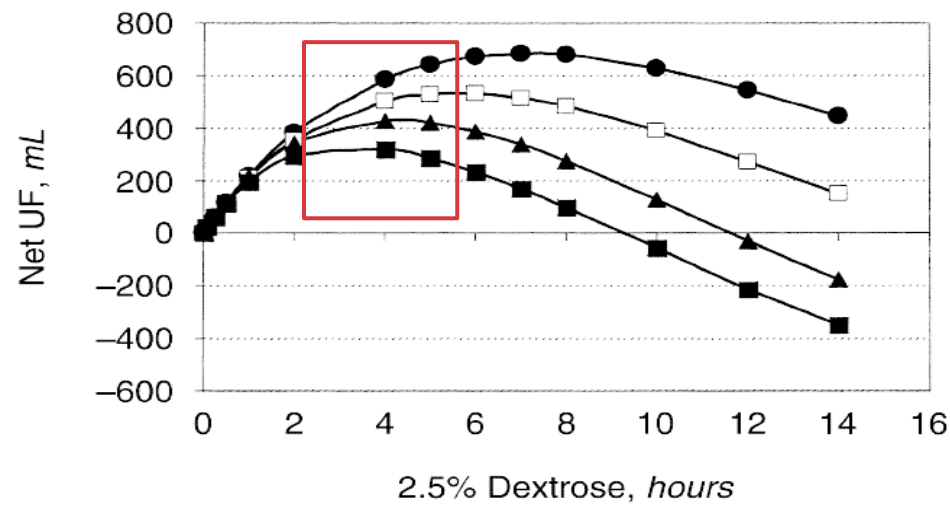
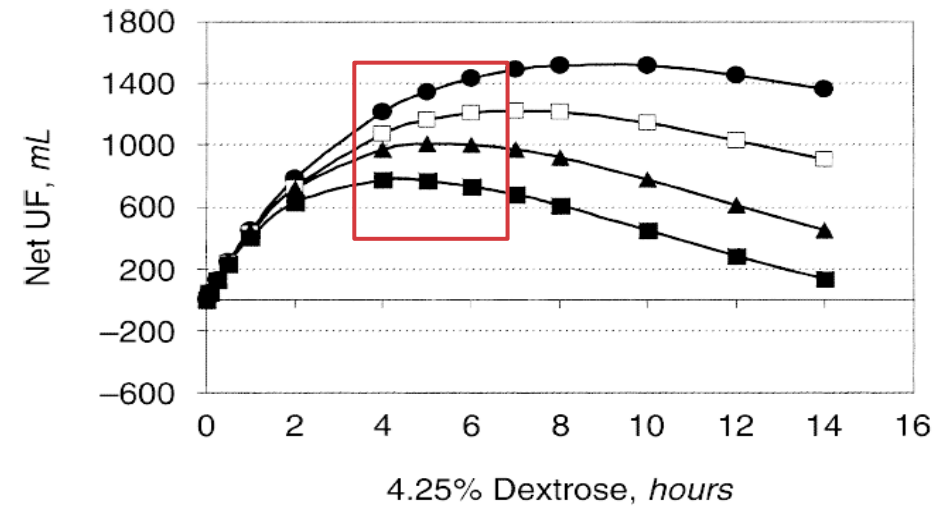
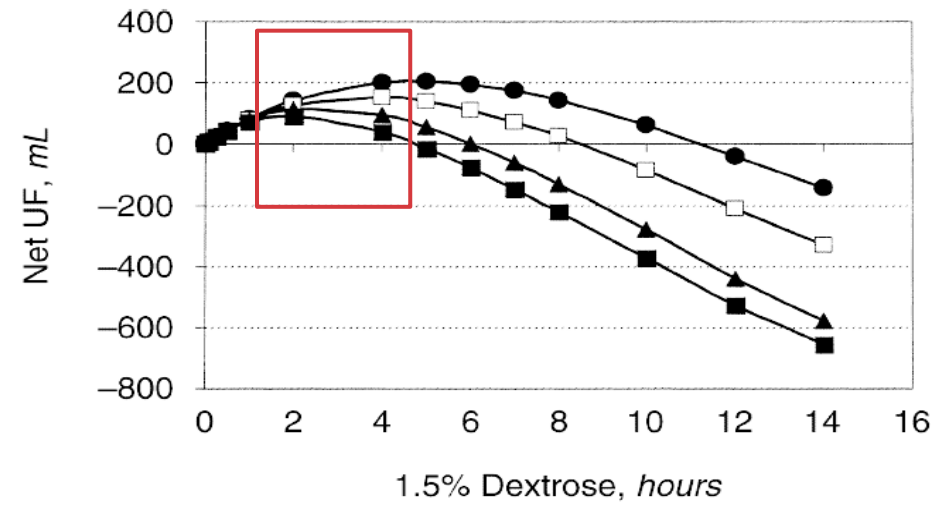
Fast vs. Slow

Fast (high) Transporter

- Reach urea/creatinine equilibrium quickly
- **Reduction in dialysate volume after ~2 hrs** (glucose absorption)
- Reduction in creatinine clearance after 4 hrs (convective creatinine re-absorption)
- Short dwells more effective
- APD often useful
- Icodextrin useful






Slow (low) Transporter

- Solute $D/P_{\text{urea/creatinine}}$ increases progressively
- UF continues **late** into dwell
- Clearance continues to increase with longer dwell times
- CAPD – (or APD (CCPD) but often less effective)

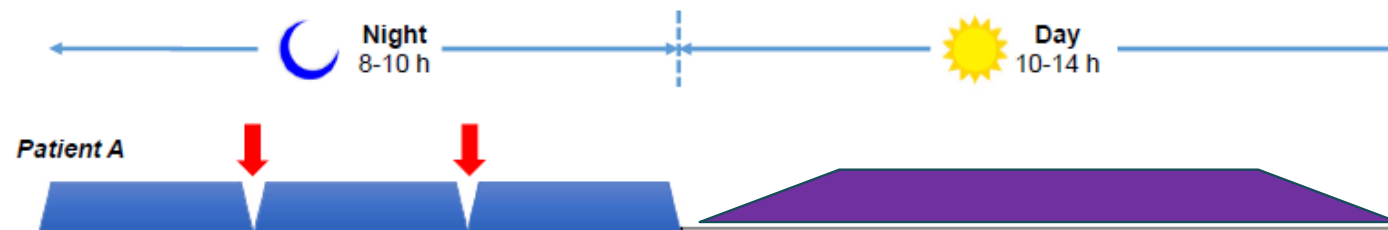


(●) low transport; (□) low average transport; (▲) high average transport; (■) high transport group.

ISPD recommendations for the evaluation of peritoneal membrane dysfunction in adults: Classification, measurement, interpretation and rationale for intervention

Johann Morelle ¹, Joanna Stachowska-Pietka ², Carl Öberg ³, Liliana Gadola⁴, Vincenzo La Milia⁵, Zanzhe Yu⁶, Mark Lambie ⁷, Rajnish Mehrotra⁸, Javier de Arteaga⁹, and Simon Davies ⁷

Guideline 2b: *Clinical implications and mitigation of fast solute transfer:* A faster PSTR is associated with lower survival on PD. (**GRADE 1A**) This risk is in part due to the lower ultrafiltration (UF) and increased net fluid reabsorption that occurs when the PSTR is above the average value. The resulting lower net UF can be avoided by shortening glucose-based exchanges, using a polyglucose solution (icodextrin), and/or prescribing higher glucose concentrations. (**GRADE 1A**) Compared to glucose, use of icodextrin can translate into improved fluid status and fewer episodes of fluid overload. (**GRADE 1A**) Use of automated PD and icodextrin may mitigate the mortality risk associated with fast PSTR. (practice point)

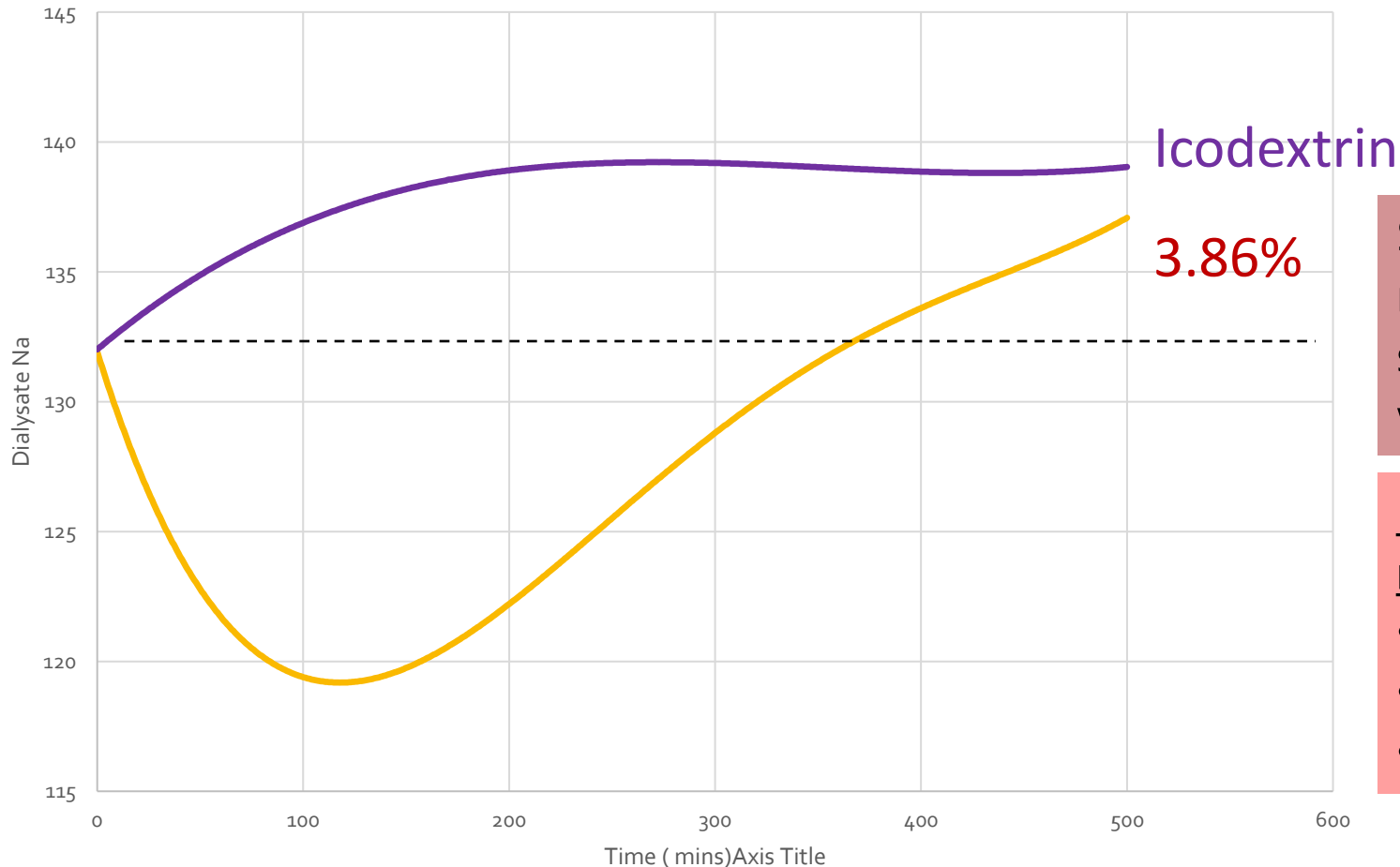


Predicted UF =

$$\begin{aligned} &\sim 300\text{mL} \times 3 \text{ (2.5\%, 3 hour dwell)} \\ &+ 600\text{mL (7.5\% 10 hour)} \\ &= \\ &1.5\text{L} \end{aligned}$$

Consider sodium sieving..

- Crystalloid osmotic gradient (glucose)
- Colloid osmotic agent (icodextrin)

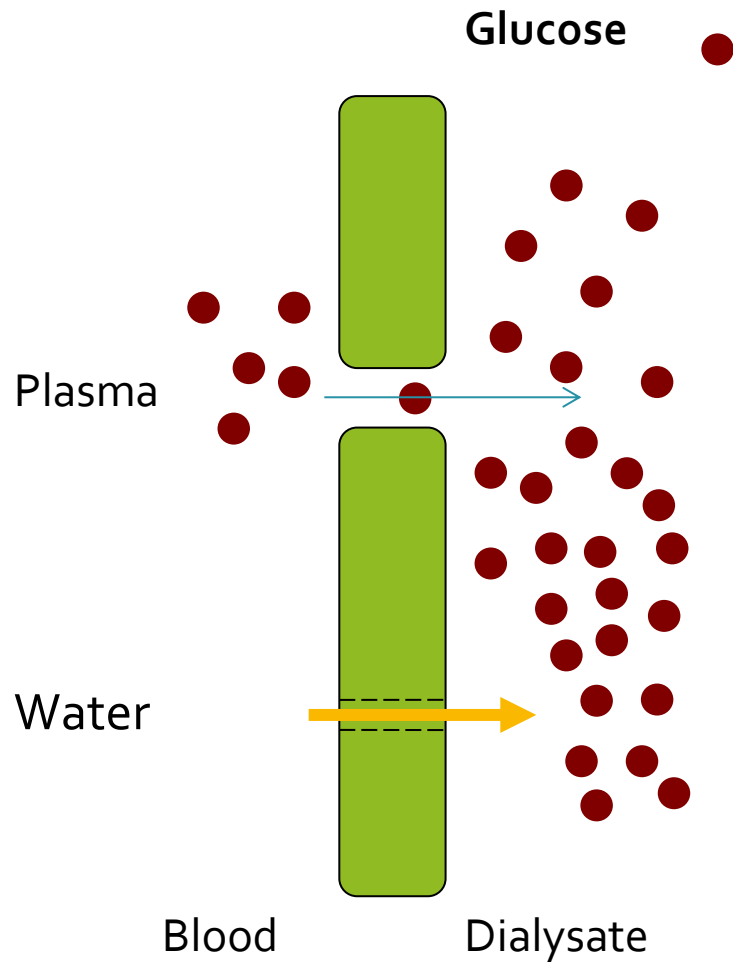


Short dwell may mean that in *less* sodium removed cf water.

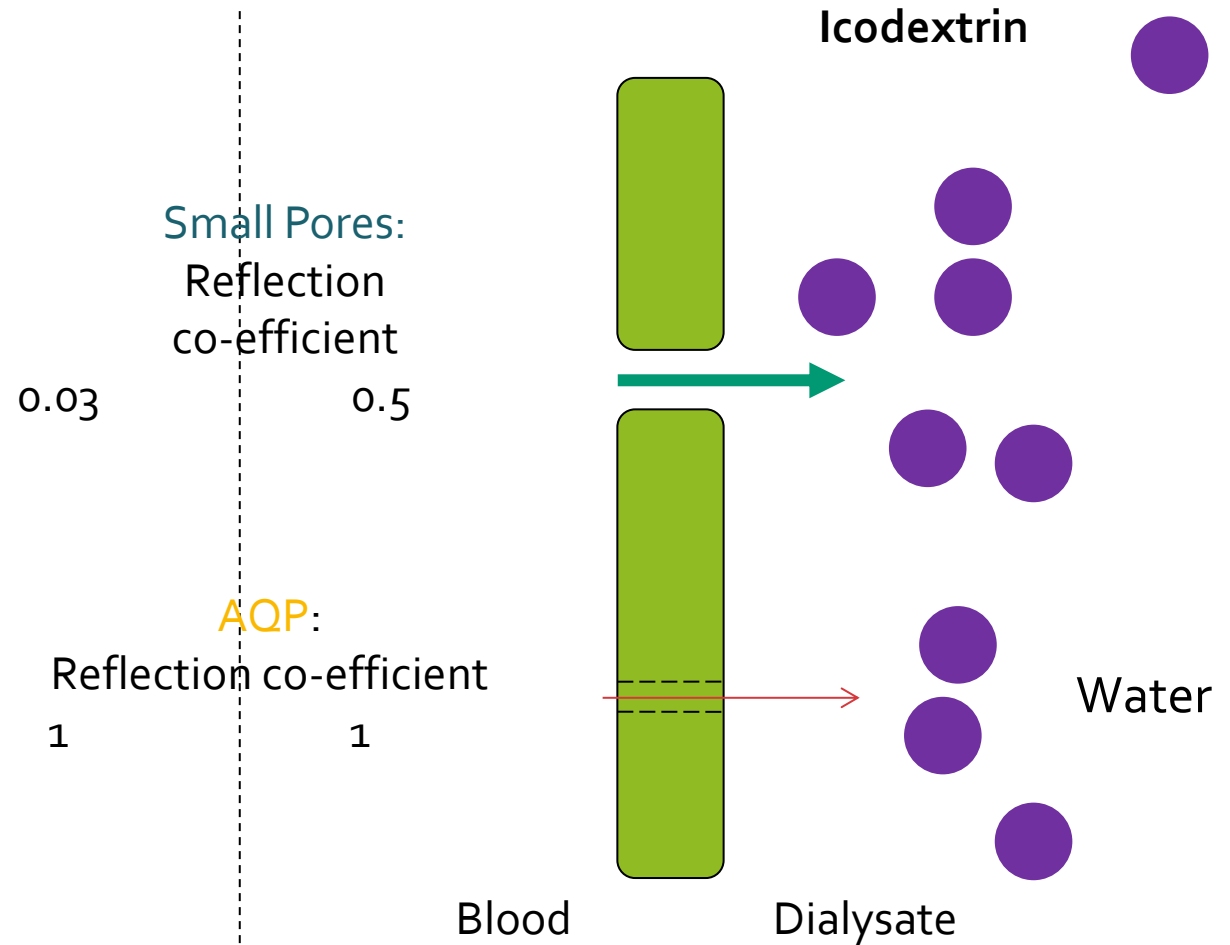
Inadequate sodium removal

- Thirst
- Hypertension
- LVH.

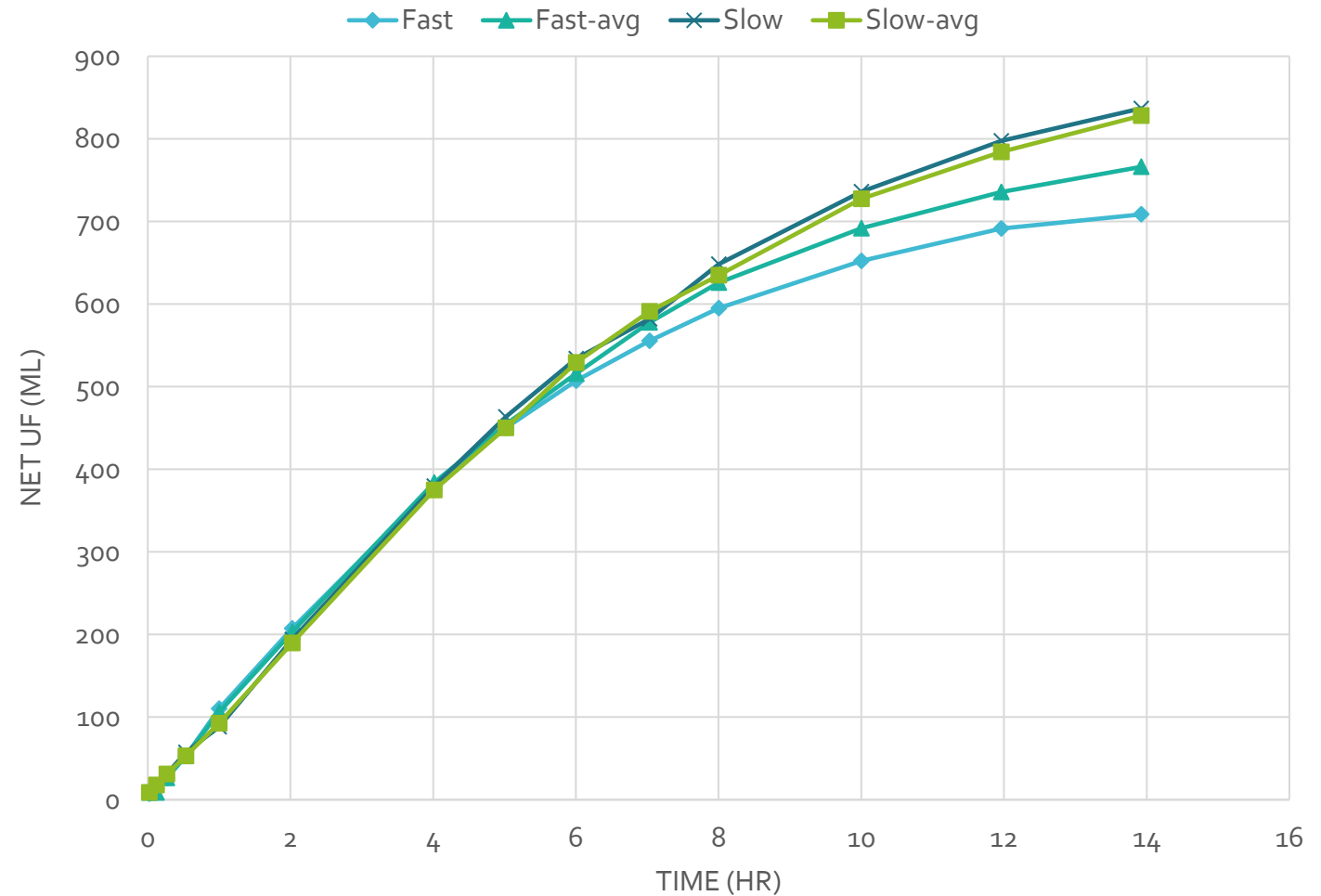
OSMOTIC pressure drives



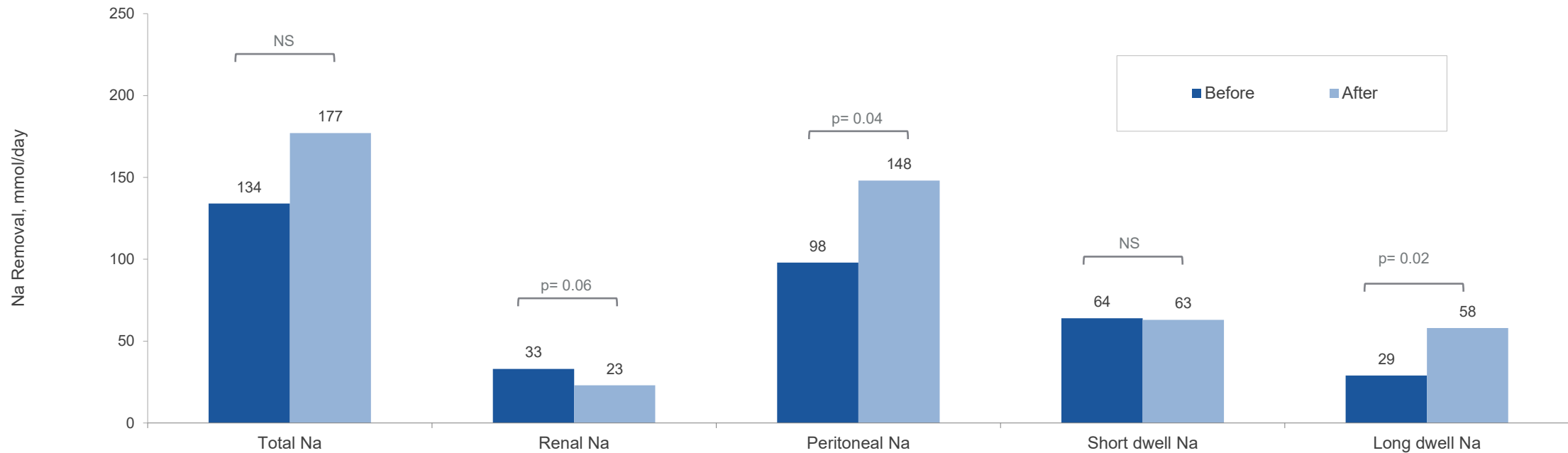
ONCOTIC pressure drives



Sustained UF with icodextrin

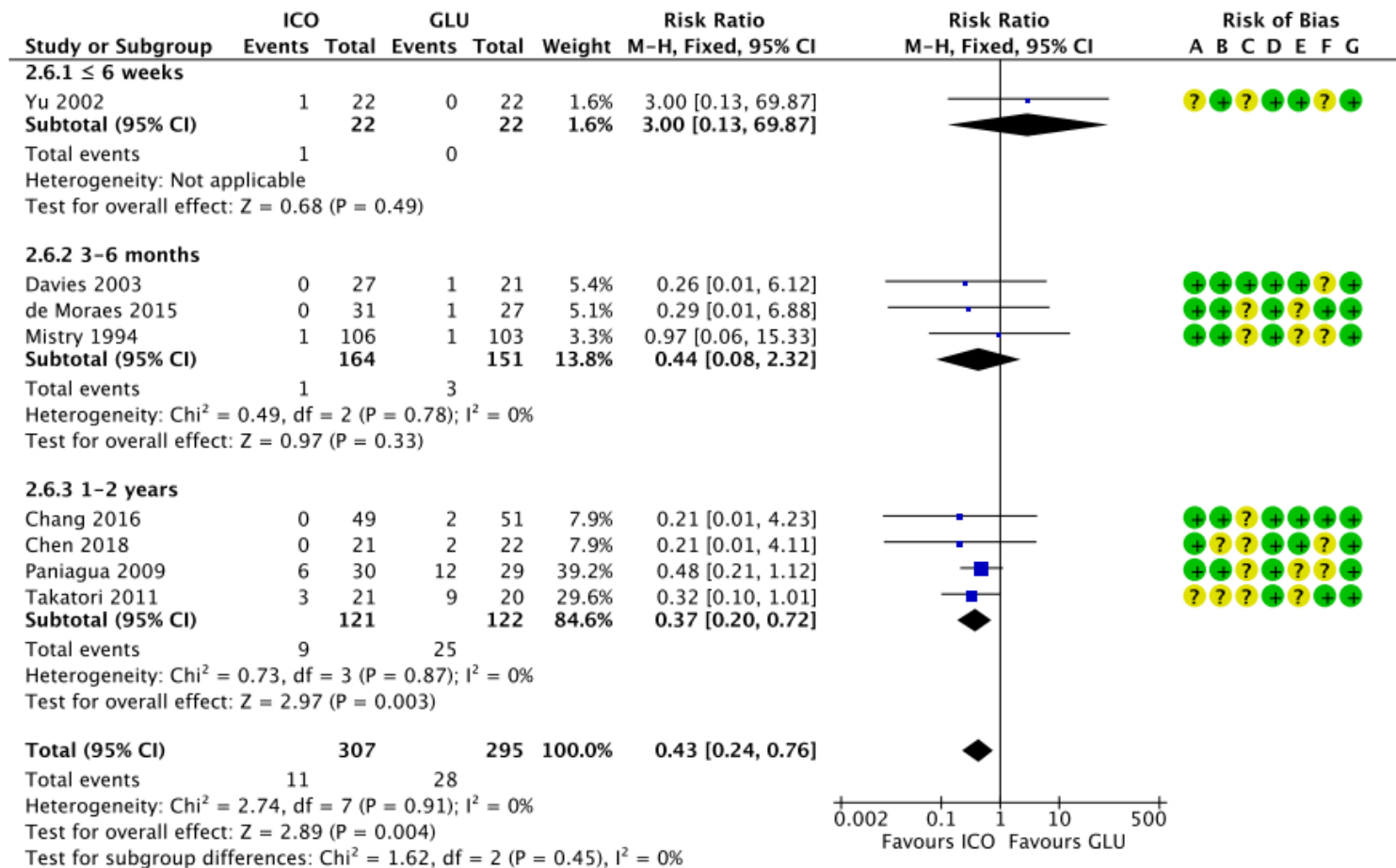


Icodextrin can remove higher levels of sodium during the long dwell when compared to glucose



Sodium Removal before and after introduction of icodextrin in 16 CAPD and APD patients.

Significantly lower risk of uncontrolled fluid overload with icodextrin use



Utility of Hybrid dialysis

- Combining PD with HD to achieve better solute and volume control
- Involves adding 1-2 HD sessions/week to a 5- to 7-day per week PD regimen.
- Majority of the published reports on hybrid dialysis comes from Japan

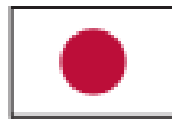
Table 4 Changes in clinical and biochemical parameters

| Author, year | Follow-up | BW | BP | Urine volume | Cr | $\beta 2m$ | Hb | D/P Cr |
|----------------------|-----------|-----------|-----------|--------------|-----------|------------|-----------|------------------------|
| Kanno, 2003 [11] | 3 months | | | | Decreased | | | |
| Kawanishi, 2006 [14] | 24 months | Decreased | Decreased | Decreased | Decreased | Unchanged | | |
| Hoshi, 2006 [15] | 36 months | | Decreased | | Unchanged | | Increased | |
| Kawanishi, 2007 [16] | 6 months | | | | | Decreased | | Unchanged or decreased |
| Matsuo, 2010 [18] | 12 months | Decreased | Decreased | Decreased | Decreased | Decreased | Increased | Decreased |
| Tanaka, 2011 [19] | 9 months | Decreased | Decreased | Unchanged | Decreased | | Increased | |
| Maruyama, 2014 [21] | 3 months | Decreased | Unchanged | Decreased | Decreased | Unchanged | Increased | Decreased |

BW body weight, *BP* blood pressure, *Cr* creatinine, $\beta 2m$ $\beta 2$ microglobulin, *Hb* hemoglobin, *D/P Cr* dialysate-to-plasma ratio of creatinine

Combination of once-weekly HD with PD is associated with lower mortality compared with PD alone: a longitudinal study

Methods



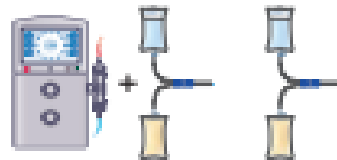
Setting

- Japanese Renal Data Registry



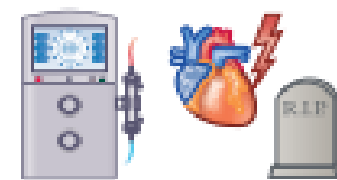
Time period

- 2010–2014



Exposure

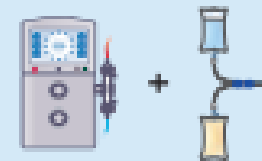
- Once-weekly HD + PD vs. PD alone



Outcomes

- Complete HD
- Mortality
- CV mortality
- CHF mortality

Results



Once-weekly HD + PD
n=608



PD alone
n=869

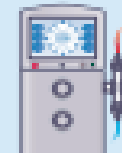


Death
n=224

In combination group:



- ↓ All-cause mortality HR 0.56 95% CI [0.42–0.75]
- ↓ CV mortality HR 0.48 95% CI [0.32–0.72]
- ↓ CHF mortality HR 0.19 95% CI [0.07–0.55]



Transition to complete HD earlier
HR 1.72 95% CI [1.45–2.03]

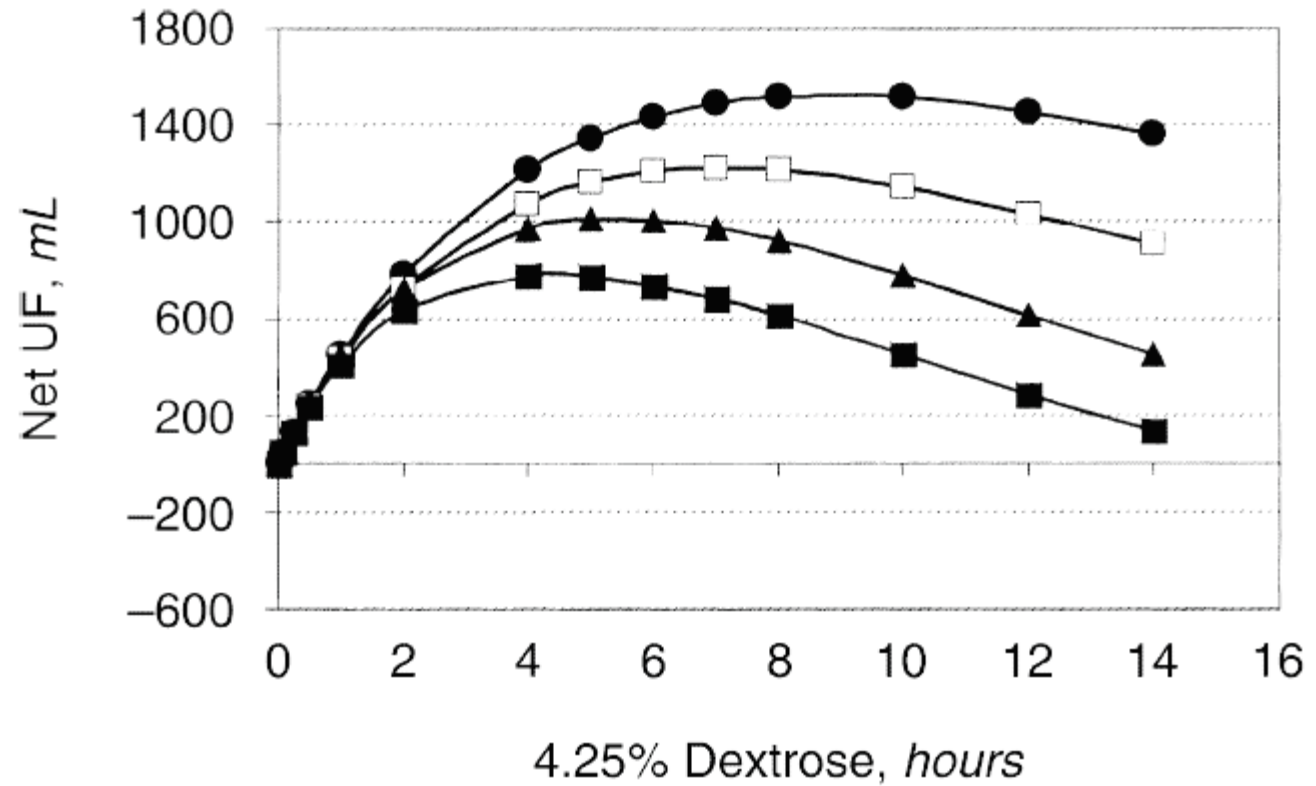
Median follow-up 2.5 years



Conclusion: Combination therapy was associated with lower all-cause mortality, CV mortality, and CHF-related mortality, but earlier transition to HD compared with PD alone.

Murashima, M. et al.
Clinical Kidney Journal (2020)
@CKJsocial

What about in
Acute Fluid
Overload – is it
time for



Take Home Message

- Fluid overload is common in PD (in both incident and prevalent patients) and an important (modifiable) cause of harm (including CVE, mortality, HD t/f)
- Target weight = symptom free + euvolaemic + normotensive
- In order to succeed in fluid management, we need to address both non-PD and PD-related issues:
 - Salt, fluid, glycaemic control
 - Maximise RKF – diuretic use
 - Optimal bowel management
 - Tailored PD prescription (and ensure adherence)

