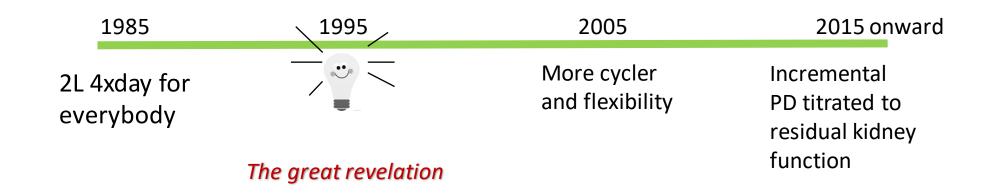
### Benefits of an Incremental PD Care Strategy Peritoneal Dialysis Workshop ICON 2022: Beyond the Call of Duty

JOANNE M. BARGMAN MD FRCPC DIRECTOR, PERITONEAL DIALYSIS PROGRAM UNIVERSITY HEALTH NETWORK, TORONTO

## My Timeline as a PD Prescriber





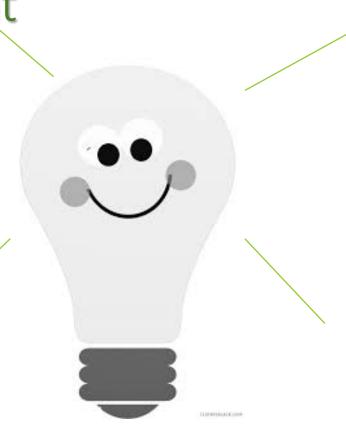
### The Great Revelation: My First Incremental PD Patient

78 year old man with advanced chronic kidney disease secondary to nephrosclerosis

GFR 8 ml/min

He and his wife would do his exchanges

"Let's start with 3 exchanges a day instead of 4 for now."



### My First Incremental PD Patient

Patient stayed on 3 exchanges/day for 5 years until he died (sudden death at home)

GFR in year 5: still 8 ml/min!

### My First Incremental PD Patient

The patient and his wife did 1 less exchange every day, compared to conventional PD

That is 365 fewer exchanges each year, X 5 years

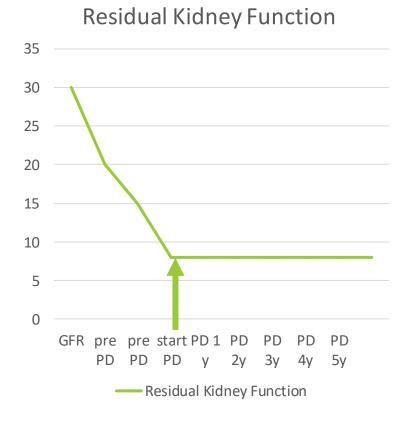
1825 saved exchanges

### My First Incremental Patient

Another Question

If the GFR declined to 8 ml/min, why didn't it continue to decline?

Why did it stabilize?



### Could PD Actually Protect the Kidneys?

Gently reduce hyperfiltration of remaining nephrons?

Removal of nephrotoxic uremic toxins?

Correction of hyperphosphatemia?

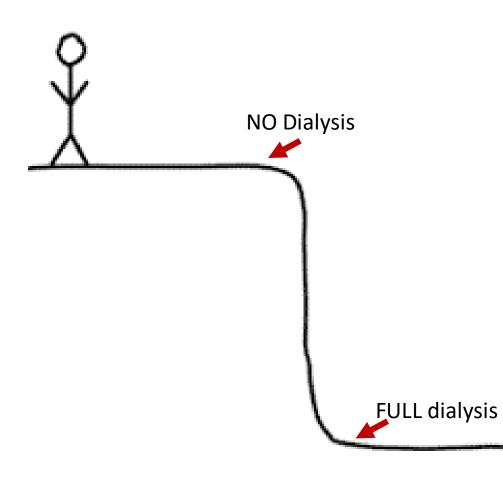
Correction of metabolic acidosis?



### What is Incremental PD?

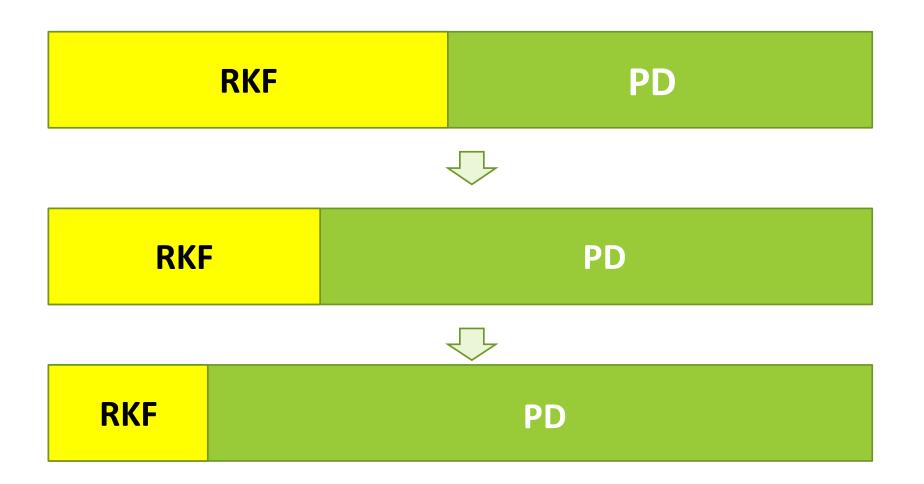
- Starting with less than the "usual" PD prescription in patients with residual kidney function (RKF)
- Increasing the dose of PD over time as the RKF declines

### From Zero to 100: Does This Make Sense?





### Incremental PD – A Schema



### Why Incremental PD? (1)

Most PD is an elective start, with "significant" kidney function (GFR 7 ml/min or more)

Small amounts of PD tend to result in improvement of symptoms

It doesn't burden the patient with the same prescription that a patient with no kidney function might need

□ It allows time for the patient to become comfortable with the therapy

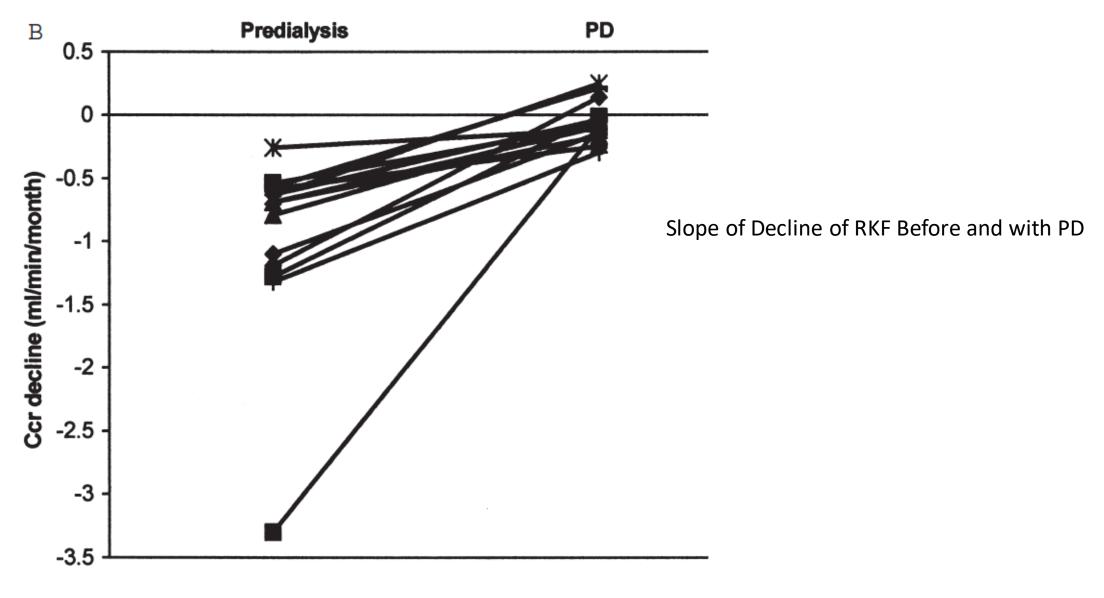
### Why Incremental PD? (2)

Less total glucose exposure

Fewer exchanges, so less risk of peritonitis

□ It saves money





Berlanga Perit Dial Int 2002

#### RATE OF DECLINE OF RESIDUAL KIDNEY FUNCTION BEFORE AND AFTER THE START OF PERITONEAL DIALYSIS

Lian He,<sup>1,2</sup> Xihui Liu,<sup>3,2</sup> Zi Li,<sup>4,2</sup> Zita Abreu,<sup>2</sup> Tushar Malavade,<sup>2</sup> Charmaine E. Lok,<sup>2</sup> and Joanne M. Bargman<sup>2</sup>

Department of Nephrology,<sup>1</sup> Peking University 3<sup>rd</sup> Hospital, Beijing, P.R. China; University Health Network,<sup>2</sup> Toronto General Hospital and the University of Toronto, Toronto, ON, Canada; Division of Nephrology,<sup>3</sup> Linyi People's Hospital, Linyi, Shandong, P.R. China; Department of Nephrology,<sup>4</sup> West China Hospital, Chengdu, Sichuan, P.R. China

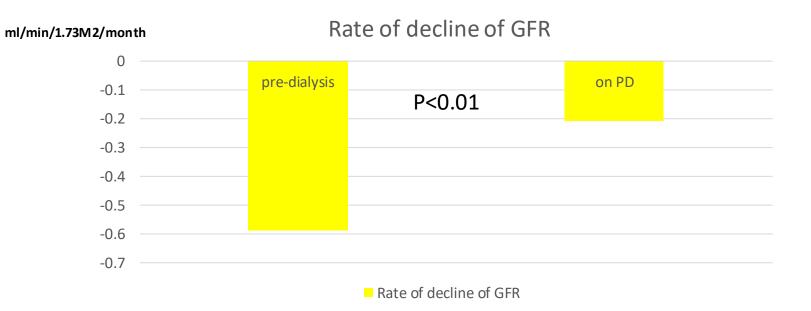
- 77 incident PD patients
- rate of decline in the predialysis period compared to rate of decline while on peritoneal dialysis
- GFR estimated in predialysis period by MDRD equation
- GFR estimated in PD by averaged 24h urine creatinine and urea clearance



#### RATE OF DECLINE OF RESIDUAL KIDNEY FUNCTION BEFORE AND AFTER THE START OF PERITONEAL DIALYSIS

Lian He,<sup>1,2</sup> Xihui Liu,<sup>3,2</sup> Zi Li,<sup>4,2</sup> Zita Abreu,<sup>2</sup> Tushar Malavade,<sup>2</sup> Charmaine E. Lok,<sup>2</sup> and Joanne M. Bargman<sup>2</sup>

Department of Nephrology,<sup>1</sup> Peking University 3<sup>rd</sup> Hospital, Beijing, P.R. China; University Health Network,<sup>2</sup> Toronto General Hospital and the University of Toronto, Toronto, ON, Canada; Division of Nephrology,<sup>3</sup> Linyi People's Hospital, Linyi, Shandong, P.R. China; Department of Nephrology,<sup>4</sup> West China Hospital, Chengdu, Sichuan, P.R. China



He Perit Dial Int 2016



	EARLY-START GROUP				LATE-START GROUP				
	n = 79				n = 72				
Overall trend over time	$-2.93 \pm 0.26$				$-3.05 \pm 0.17$				
Trend during the pre- and post-dialysis initiation	PRE	POST	CHANGE		PRE	POST	CH		
periods			Value (95% CI)	P value			Value (95%		
Unadjusted model	-4.55 ± 0.68	$-2.71 \pm 0.28$	+1.84 (0.43-	< 0.001	$-4.08 \pm 0.39$	-2.50 ± 0.19	+1.58 (0.6		
	('	(′	3.25)	<u> </u>	<u> </u>	<u> </u>	2.46)		
Exploratory model*	$-4.68 \pm 0.71$	-2.71 ± 0.28	+1.97 (0.50-	0.008	$-4.07 \pm 0.39$	$-2.49 \pm 0.19$	+1.58 (0.7		
	('	(′	3.44)	//	('	(′	2.46)		

Table 4. Trend of glomerular filtration rate over time (in mL/min/1.73m<sup>2</sup>/yr) in the early and late dialysis start groups.

\*Adjusted for patients' characteristics at enrollment: age, sex, ethnicity (Caucasian vs non-Caucasian), initial dialysis dose (incremental vs full), presence of mellitus and history of cardiovascular disease.





#### **Original article**

### 

Mercè Borràs Sans\*, Andrea Chacón Camacho, Carla Cerdá Vilaplana, Ana Usón Nuño, Elvira Fernández

Servicio de Nefrología, Hospital Universitari Arnau de Vilanova, Lérida, Spain

46 patients receiving 3 or fewer exchanges/day

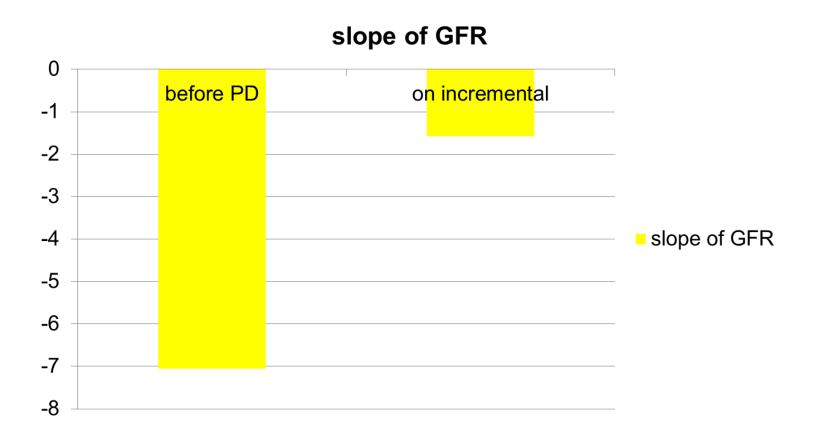
24 months

many were transplanted before moving to standard PD

good outcomes



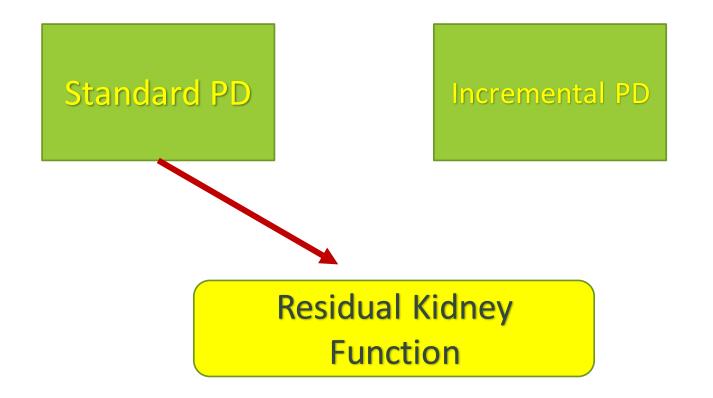
### Incremental PD, Like Full-Dose PD, May Also be Kidney Protective



Barras Sans Nefrologia 2016

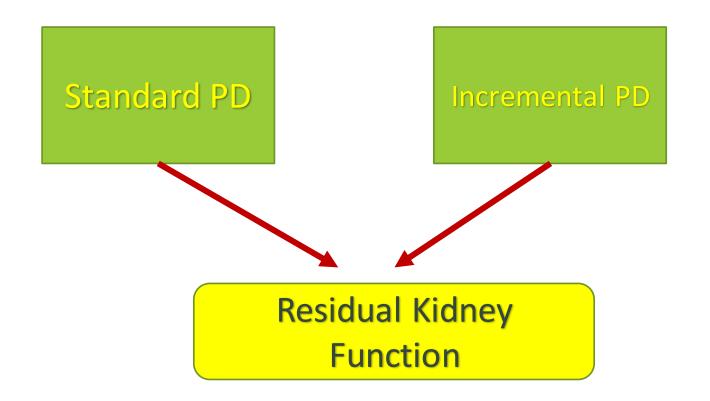


### In Fact, Could Incremental PD be Even *More* Protective of Kidney Function?



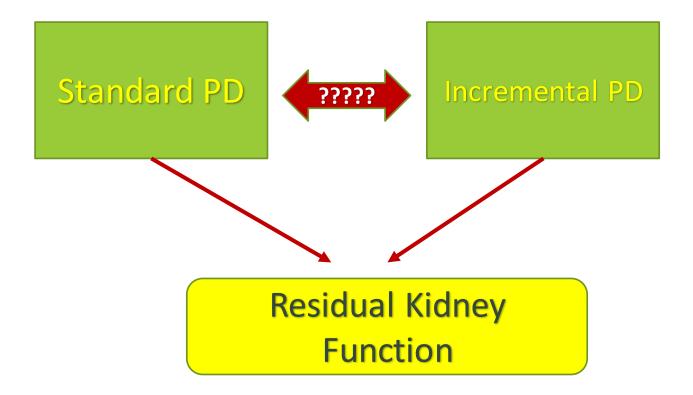


# Could Incremental PD be Even More Protective of Kidney Function?





# Could Incremental PD be Even More Protective of Kidney Function?





## Outcome of Incremental PD

Single center, retrospective study over 16 years

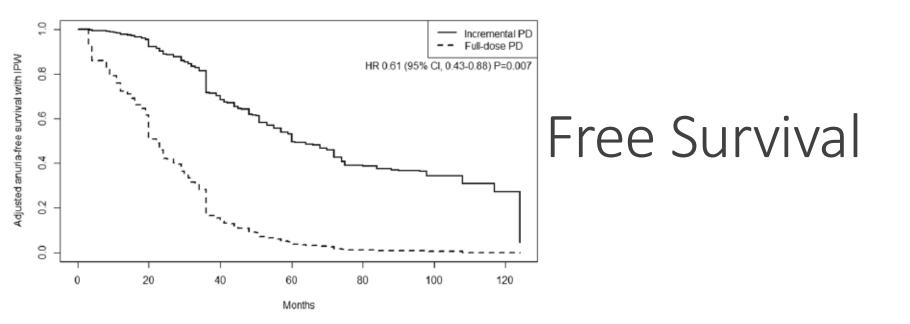
Patients commenced PD between 2007 and 2015

Propensity scores, IPW adjustments +++

Incremental PD defined as 1-2 exchanges/day, 7 days a week

The incremental group had lower risk of developing anuria (HR 0.61, 95% CI .43-.88)

Technique survival, patient survival, peritonitis no different between the two groups



**Incremental PD** 

**Full-Dose PD** 

Lee Sci Reports 2019

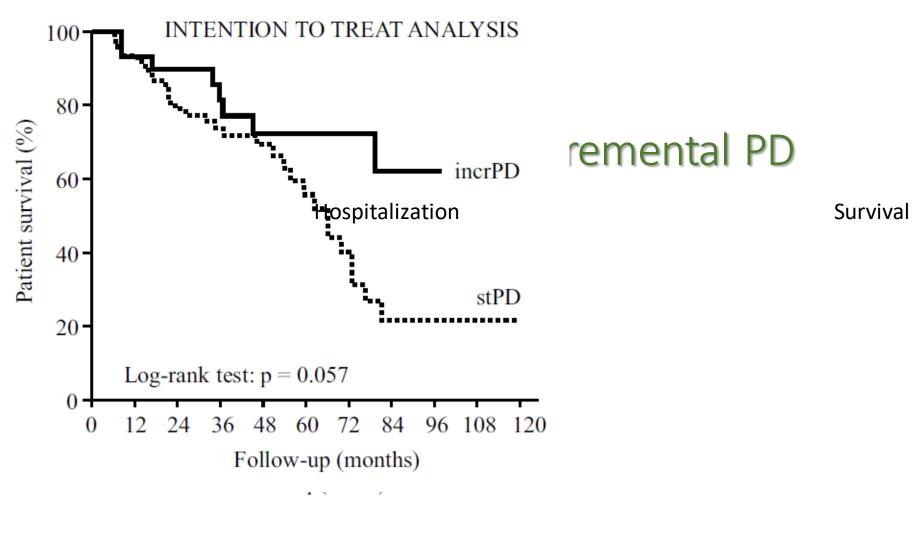
	incrPD	stPD	р
Number of patients	29	76	
Male gender	13 (55%)	50 (66%)	0.611
Age (years)	$63 \pm 12$	59±18	0.200
Weight (Kg)	$63.4 \pm 10.2$	62.8±16.7	0.837
BMI (Kg/m <sup>2</sup> )	24.3±3.9	$23.3 \pm 3.7$	0.130
RRF (ml/min/1.73 m <sup>2</sup> BSA)	$5.74 \pm 1.34$	$5.42 \pm 1.75$	0.381
D/P creatinine 4th hour	$0.63 \pm 0.14$	$0.62 \pm 0.11$	0.426
-23 palients		וונמו (ודע) ו	เคริแบรเ

 Table 1 Baseline data of the two groups: incrPD and stPD

•76 patients on standard PD prescription (stPD)

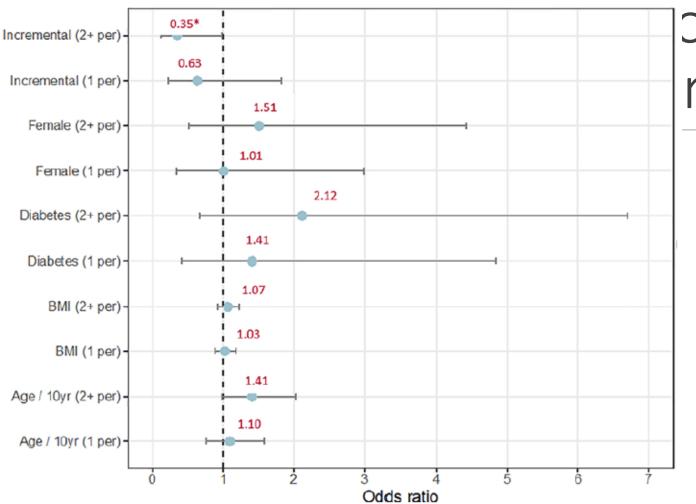
Median duration of iPD was 17 months

Same baseline GFR in the two groups



#### Figure 2

Predictors of peritonitis. Multivariate logistic regression analysis



# on is Associated

\*\*

Santos Port J Nephrol Hypert 2021



### Table III

Predictors of drop out to hemodialysis. Multivariate analysis

	OR	95% CI	P value	
Age (per decade)	0.97	0.75-1.26	0.814	
DM	1.52	0.69-3.34	0.303	
Albumin <mark>(</mark> g/L)	0.89	0.38-2.11	0.798	
Body mass index (per Kg/m <sup>2</sup> )	1.0	0.89-1.13	0.999	
Peritonitis in the first year	1.83	0.79-4.27	0.158	
PD start decade	1.74	0.77-3.91	0.182	
GFR (mL/min)	0.94	0.82-1.08	0.401	
Incremental approach	0.41	0.19-0.92	0.030*	

# ion is Associated ique Survival

\*

transfer to hemodialysis

### How To Prescribe Incremental PD

CAPD

### One exchange overnight

- if using a 1.5% solution, there will likely be absorption, so it depends on the fluid status of the patient
- can use 2.5% solution or icodextrin if available
- the icodextrin will usually result in ultrafiltration



### How To Prescribe Incremental PD

CAPD

- 2 exchanges, 4h each during the day, night dry
  - this works well in patients with RKF
  - good incremental regimen for those who don't like fluid in the abdomen overnight



### How NOT To Prescribe Incremental PD

CAPD

2 exchanges, 12 hours each may not be successful

• there will likely be fluid absorption that can result in volume overload



### How To Prescribe Incremental PD

APD

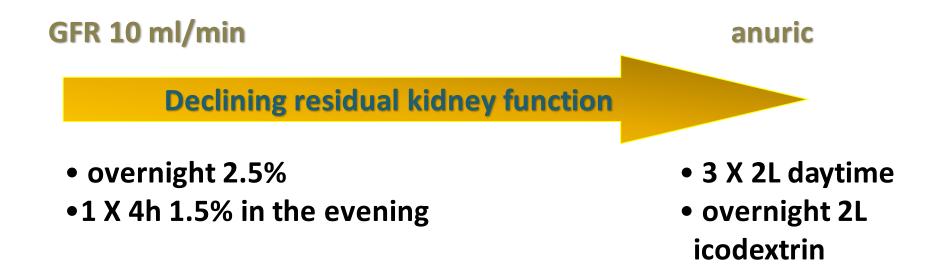
Night cycles, day dry is a great regimen (NIPD)

don't have to worry about fluid absorption during the long day dwell

• Example: 3 X 1.5 L exchanges over 8 hours

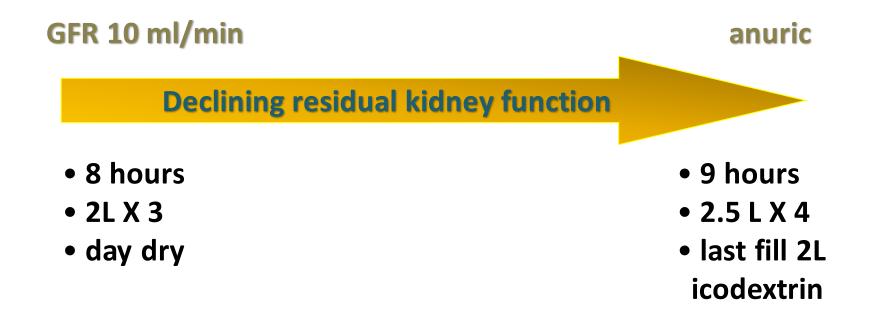


## Titrating the Dose of Dialysis to Residual Kidney Function (CAPD)





## Titrating the Dose of Dialysis to Residual Kidney Function (APD)





### The Volume Can Also be Increased Incrementally

- No need for a full 2 or 2.5L dwell volume at the outset
- Allow time for adjustment to the sensation
- It takes 9 months to grow a baby for a reason





## The <u>Best</u> Options for Increasing the Dose of Dialysis in the Incremental APD Patient

Add a day dwell

Increase the volume of the night dwells



## The <u>Worst</u> Options for Increasing the Dose of Dialysis in the Incremental APD Patient

Increase the number of exchanges overnight

 Increase the time on the cycler, especially if the patient has a day dwell

# An <u>Ineffective</u> Way to Increase the Dose of Peritoneal Dialysis





## A *Really* Ineffective Way to Increase the Dose of Peritoneal Dialysis

9 hours PD	15 hour day dwell
10 hours PD	14 hour day dwell





**Standard PD** 



### FY 2019/20 Top Performance Award

This certificate is awarded to

## University Health Network

For achieving the highest Home Dialysis Prevalence in Ontario



Congratulations to all staff and thank you for your continued contribution and commitment towards improving the lives of Ontario renal patients

Elister

**Garth Matheson** Vice President (Temporary) Ontario Renal Network

Cover State

**Dr. Peter Blake** Provincial Medical Director Ontario Renal Network





# What Are The Objections to Incremental PD?

In many units there is a reluctance to use incremental PD, because there is no "evidence" that it works and is safe

 Observational studies are confounded by indication: stable patients with good RKF would be more likely to receive an incremental prescription, but this group would do better in any case

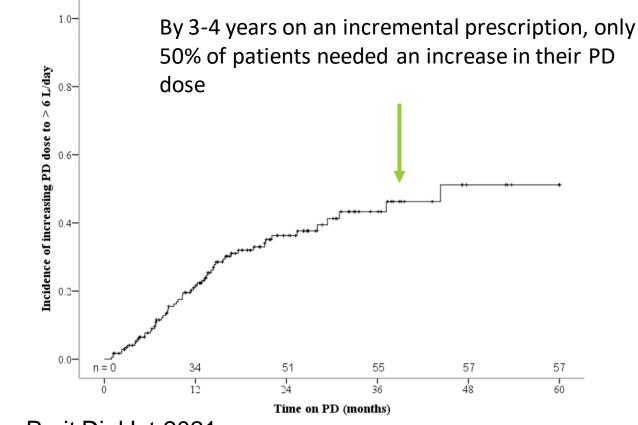


#### "The Patient Will Refuse to Increase the Prescription Once the Kidney Function Declines"

- I haven't found this to be a problem
- The process has to be explained and reexplained to the patient
- Actually, transitioning from night cycler/day dry to night cycler day dwell is not usually a big disruption to life



# Cumulative Incidence of Increasing the PD Dose to > 6L/Day: Our Experience



Yan Abreu and Bargman Perit Dial Int 2021

10%

#### "If You Have to Get to Full Dose Eventually you Might as Well Start Off with It"

- in our experience many patients keep their RKF until death or kidney transplant or transition to HD
- Would you proceed directly to an above-knee amputation instead of a below-knee amputation if the patient may need it in the future?



#### "The Patient May Lose RKF Suddenly, and They Will Quickly Become Under-dialysed"

 ambulatory PD patients don't suddenly lose their RKF

 sudden loss of RKF is usually in the context of an intercurrent event (CHF, radiocontrast, sepsis) which, hopefully you will be aware of and can make adjustments to the PD prescription

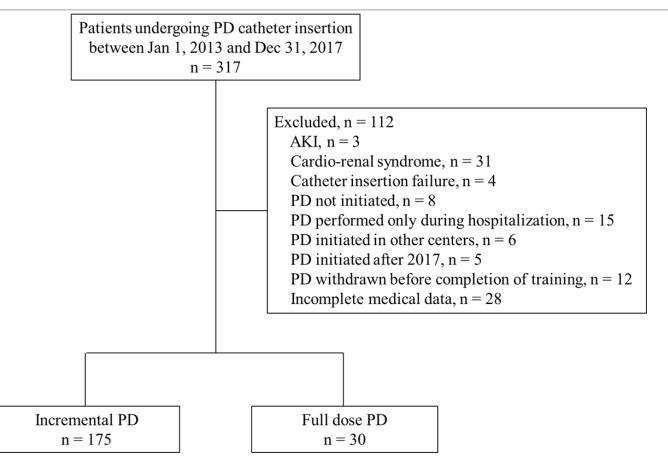


#### "You Have to Carefully Monitor the Residual Kidney Function with 24h Urine Collections"

- Worth repeating: ambulatory PD patients don't suddenly lose their RKF without other stuff happening
- We request a 24h urine for volume/urea/creatinine every 2 to 4 months
- If the patient "forgets": if the serum creatinine is unchanged, and the prescription is unchanged, then we presume the kidney creatinine clearance is unchanged



#### Walking the Walk: The University Health Network Experience Yan, Abreu and Bargman Perit Dial Int 2021





#### "There are No Randomized Trials to Prove that This Approach is Safe"

#### true

- •however, I think that most people would take 5 ml/min of KIDNEY clearance any day compared to 5 ml/min of dialytic (HD or PD) clearance
- using kidney Kt/V urea grossly underestimates what the kidney function is doing (volume, larger molecular weight toxin removal)





#### Incremental Dialysis: Summary

Someone with <u>some</u> kidney function does not need the same dialysis prescription as someone with <u>no</u> kidney function

- Incremental PD may preserve kidney function, allowing incremental dialysis to continue
- There are many potential benefits, especially for quality of life and illness intrusion

