WESTERN CANADA DDD DAYS



February 7th-9th, 2013

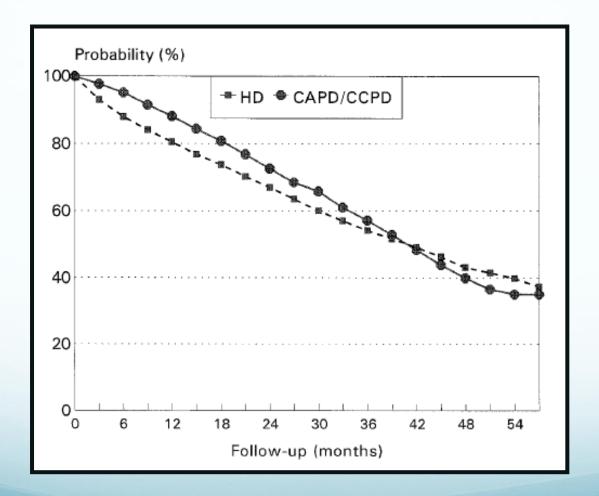
PD Outcomes: Do Patient Socio-Demographics Matter?

Outline of Presentation

- Does the relative risk for death with PD, compared to that seen with HD, change over time?
 - Is PD better treatment early in the course of ESRD but an inferior long-term therapy?

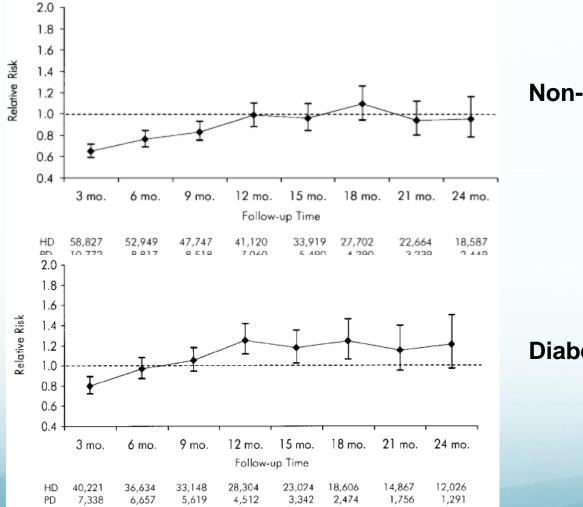
- PD outcomes in selected sub-groups:
 - Regional variation
 - Racial/ethnic minorities
 - Poverty/Educational attainment
 - Rural communities or with greater distance from dialysis facilities?

PD vs. HD Death Risk: Change over Time?



Fenton et al, Am J Kidney Dis 1997, 30: 334-42

PD vs. HD Death Risk: Change over Time?



Non-diabetics

Diabetics

Collins et al, Am J Kidney Dis 1999; 34: 1065-74

Conventional Wisdom

- Patients treated with PD have an early "survival advantage":
 - Residual renal function is preserved better with PD
 - So, this early survival advantage is a direct benefit of being treated with PD

- However, over time the death risk of PD patients goes up:
 - This happens because patients lose residual renal function
 - Without residual renal function, PD alone is inadequate for removing enough urea or fluid
 - Moreover, glucose in PD fluids is really bad, damages the peritoneum, and makes volume control quite challenging

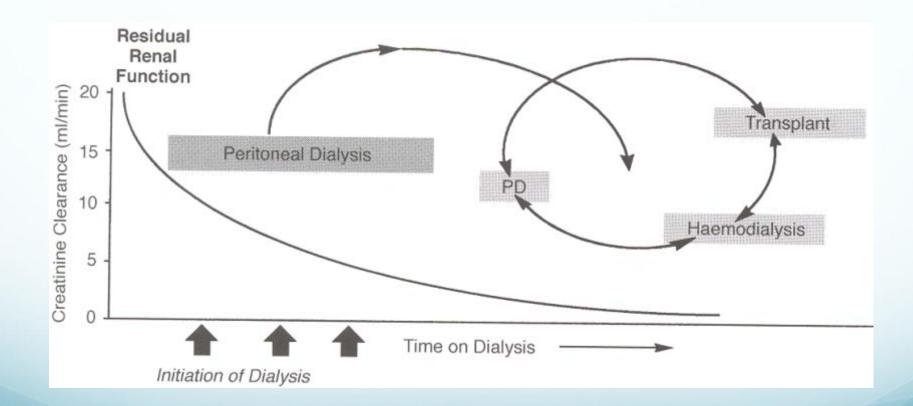
PD Needs Residual Renal Function

Variable	RR	95% CI
Age	1.02	1.01-1.041
CVD	2.42	1.50-3.90
Diabetes mellitus	1.25	0.77-2.04
Serum Albumin	0.96	0.91-1.00
LA Transport	1.66	0.38-7.22
HA Transport	2.33	0.55-9.80
High Transport	2.01	0.43-9.36
SGA	0.74	0.65-0.84
Ccr (perit 5 L/wk/1.73 m ² ↑)	1.00	0.90-1.11
GFR (5 L/wk/1.73 m ² ↑)	0.88	0.83-0.94

Bargman et al, J Am Soc Nephrol 2001; 2158-62

Conventional Wisdom

Van Biesen and Lameire



Start more patients with PD but make sure you transition them to HD in a timely manner, particularly if they lose residual renal function

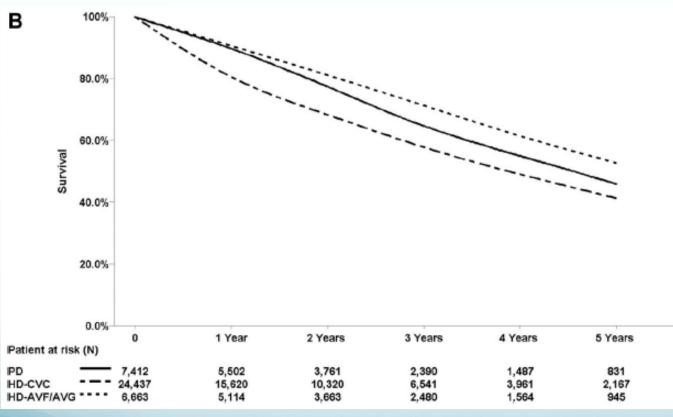
Is the early lower death risk of patients who start PD an effect of PD?

Early "Survival Advantage" with PD?

Better preservation of residual renal function with PD?

- Many patients start dialysis sub-optimally (hospitalized, without a permanent access for the dialysis therapy of their choice):
 - These are the patients with poorer outcomes
 - Most of them start hemodialysis with central venous catheters

Early Survival Advantage with PD?



CORR Registry Data

79% of HD patients started with CVC

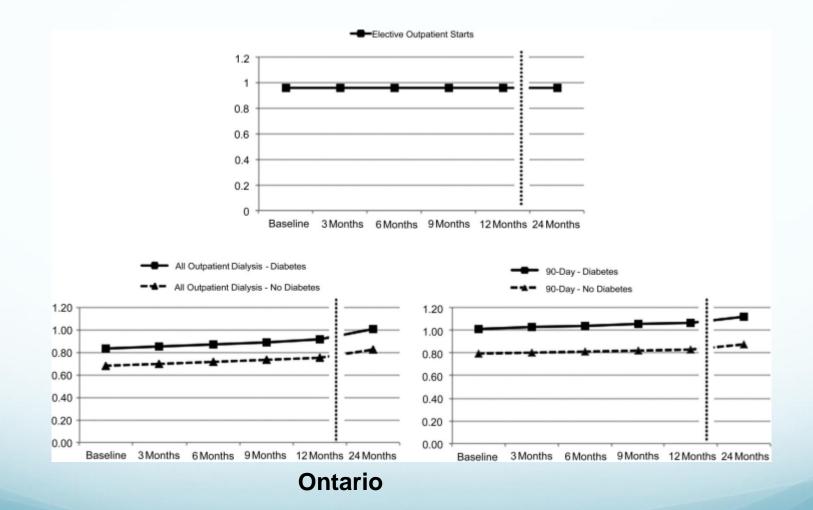
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PD survival better only with those who started with CVC and not AVF/AVG

Perl et al, J Am Soc Nephrol 2011; 22: 1113-21

Early Survival Advantage with PD?



Quinn et al, J Am Soc Nephrol 2011; 22: 1534-42

An Alternative Wisdom?

- Early survival of PD patients looks better only because patients that start HD do so much worse:
 - Most patients with no pre-dialysis nephrology care are treated with HD
 - Virtually all these patients are treated with central venous catheters that increases their death risk
 - It is not HD that is bad, but how patients are treated when they start with the therapy
- Death risk of PD patients does not really go up over time:
 - It just seems that way as sicker patients who start with HD earlier
- The "change in relative risk over time" is not the direct effect of dialysis therapies but a result of differences in patients who are treated with these therapies

• Where does residual renal function fit into this wisdom?

Is residual renal function more important for PD?

RRF is Important for PD But it is also important for HD

Table 4. Multivariate Cox regression models with residual renal Kt/V_{urea}, delivered Kt/V_{urea}, and the net fluid balance as independent predictors of survival^a

	Adjusted RR			
	RR	95% CI	P Value	
rKt/V _{urea} (/wk)			< 0.0001	
0	17.66	4.98 to 62.61		
>0 to 0.84	1.67	1.06 to 2.64		
>0.84	1.0 ref			
sp-dKt/V _{urea} (/wk)				
if rKt/V _{urea} = 0	0.54	0.40 to 0.72	< 0.0001	
if rKt/V _{urea} > 0	0.90	0.72 to 1.12	0.3395	
Net fluid balance (ml/wk) ^b			0.0021	
≤-300	2.17	1.46 to 3.22		
>-300 to ≤-50	1.24	0.82 to 1.86		
>−50 to ≤67	1.00 ref			
>67 to ≤300	1.31	0.85 to 2.00		
>300	1.35	0.88 to 2.07		

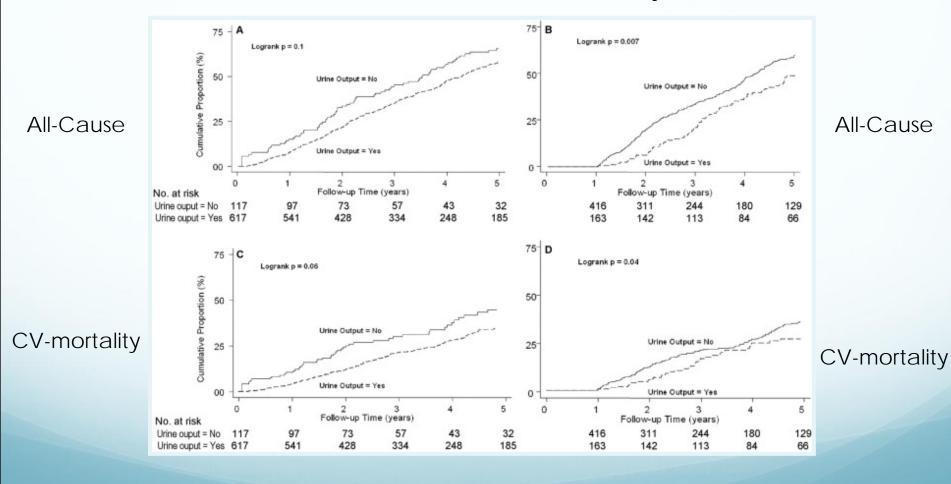
^a The shown RR are adjusted for age, Davies' comorbidity score, primary kidney disease, SGA, and BMI. The rKt/V_{urea}, the dKt/ $V_{\rm urea}$, and the net fluid balance were entered as time-dependent variables.

Termorshuizen et al, J Am Soc Nephrol 2004; 15: 1061-70

RRF Is Important for HD

Baseline U.O. > 250 ml

U.O. at 1-year > 250 ml



Shafi et al, Am J Kidney Dis 2010; 56: 348-58

Residual Renal Function in Dialysis Patients

• Is important for both PD and HD patients

- Loss of residual renal function makes it challenging to manage both HD and PD patients and increases the death risk of both:
 - Don't overestimate the value of ultrafiltration of large volumes of fluid with each HD session
- It is questionable that the apparent "change in relative risk over time" is related to differential importance of residual renal function

PD can be successfully done even in anuric patients

Region	Author	N	Mean Age	% diabetics	CAPD/APD	Two-year Survival
Hong Kong	Szeto, '01	140	53 y	26%	140/0	69%
	Lo, '05	150	58 y	28%	149/1	89%
				27%	289/1	79%
Europe	Brown, '03	177	54 y	42%	0/177	78%
	Jansen, '05	130	53 Y	12%	102/28	67%
				29%	102/205	73%
United States	Fried, '08	1428	54 y	42%	?	60%

Only somewhat worse than PD patients with residual renal function

Is the long-term survival of PD patients as good as that with HD?

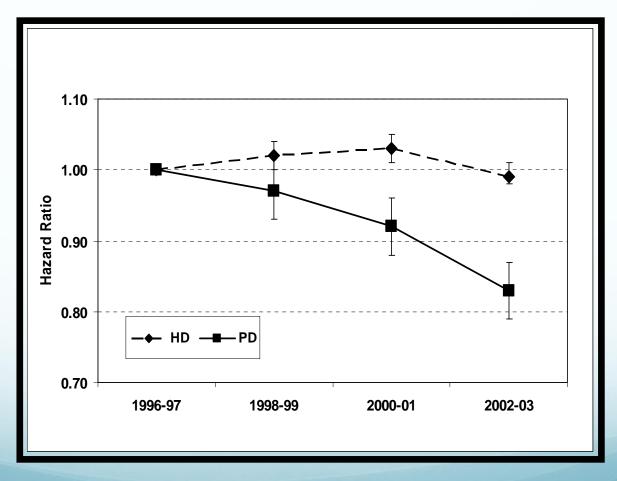
Greater Improvements in Survival of PD Patients

	1996-'97	1998-'99	1999-2000	2000-'01
Death – HD	22.3%	23.0%	23.5%	22.8%
PD	17.2%	16.6%	15.2%	14.0%
Transfer – HD PD	1.8%	1.5%	1.5%	1.4%
	12.7%	12.4%	12.5%	12.6%
Transplant – HD	2.8%	2.8%	2.8%	2.5%
PD	7.3%	7.5%	7.6%	7.2%

Outcomes in First 12 months

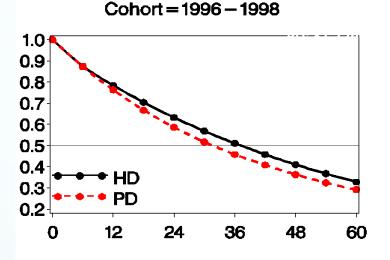
Mehrotra et al, J Am Soc Nephrol 2007; 18: 2781-8

Greater Improvement in PD Outcomes



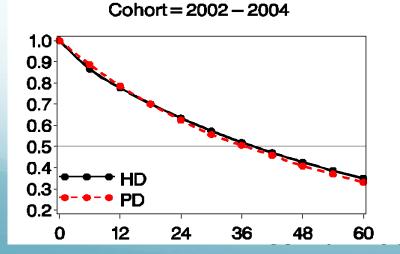
Mehrotra et al, J Am Soc Nephrol 2007; 18: 2781-8

Equal Five-Year Survival with HD and PD



1.0⁻ 0.9 0.8 0.7 0.6 0.5 0.4 0.3 PN 0.2 24 12 36 48 60 0

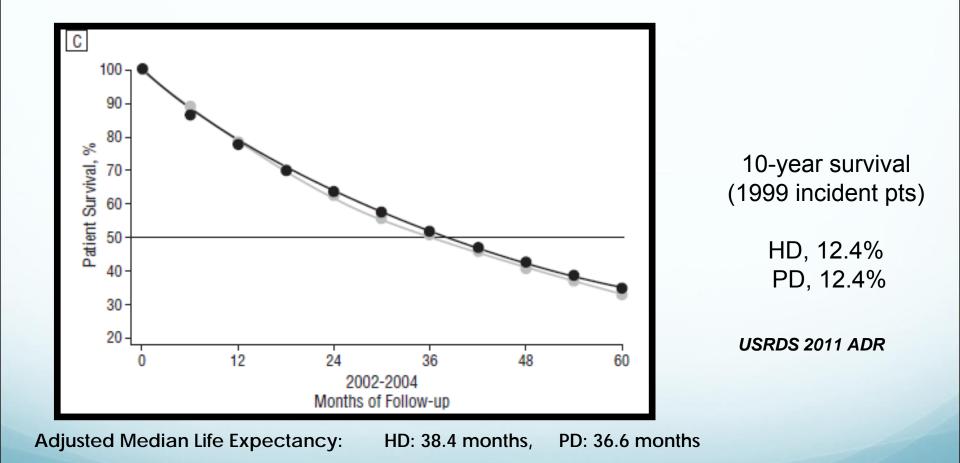
Cohort = 1999 - 2001





Mehrotra et al, Arch Intern Med 2011; 171: 110-8

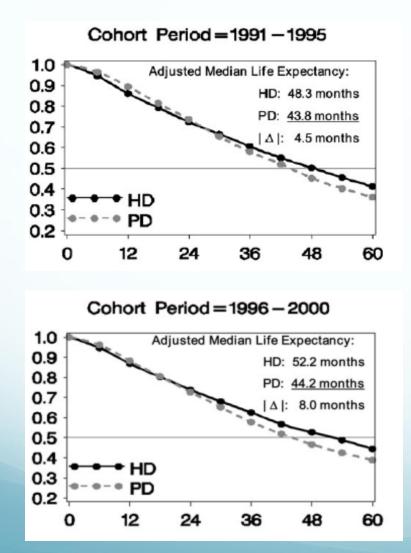
And In the Longer Term



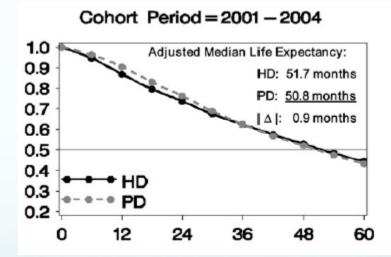
Mehrotra et al, Arch Intern Med 2011; 171: 110-8

Survival with HD and PD is equivalent even when a lot more patients use PD

Similar Survival Canada

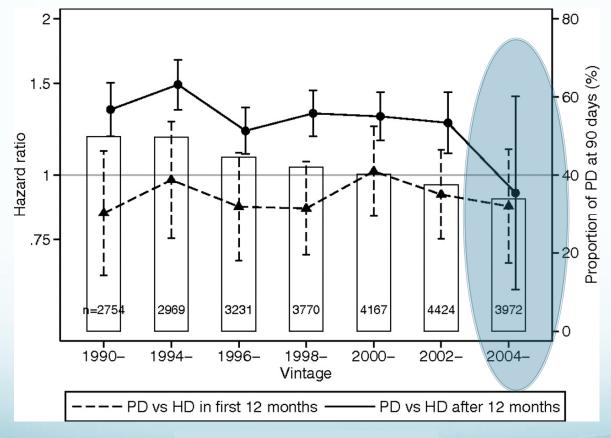


PD use, 18%



Yeates et al, Nephrol Dial Transplant, (epub)

Similar Survival Australia-New Zealand

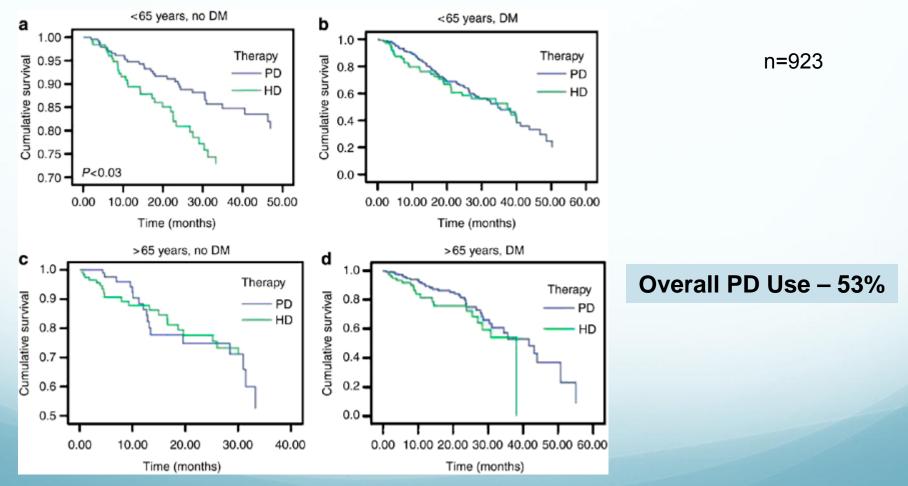


n=25,287

Overall PD Use – 25%

McDonald et al, J Am Soc Nephrol 2009; 20: 155-163

Similar Survival Colombia



Sanabria et al, Kidney Int suppl 2008; 73: S165-72

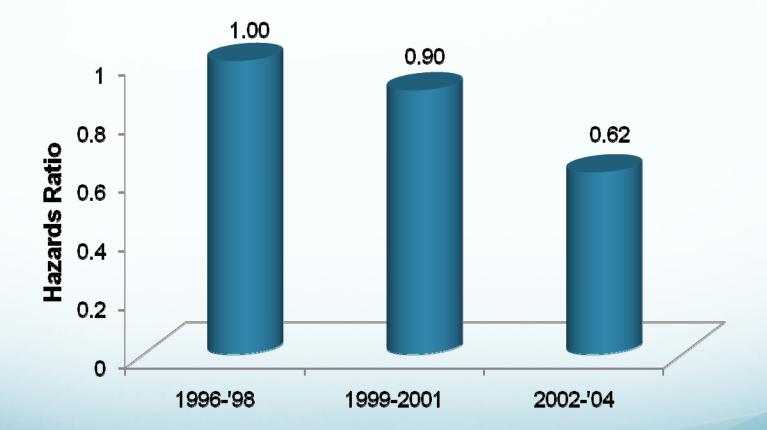
What Have I Said So Far?

- The dialysis therapy that the patient is treated with does not specifically determine how a patient does:
 - Neither early in the course or over the long-term
 - Patients do equally well in the short-term or long-term with PD or HD

- Residual kidney function is important for both hemodialysis and peritoneal dialysis:
 - That it matters only for PD or PD cannot be done well without it is misleading

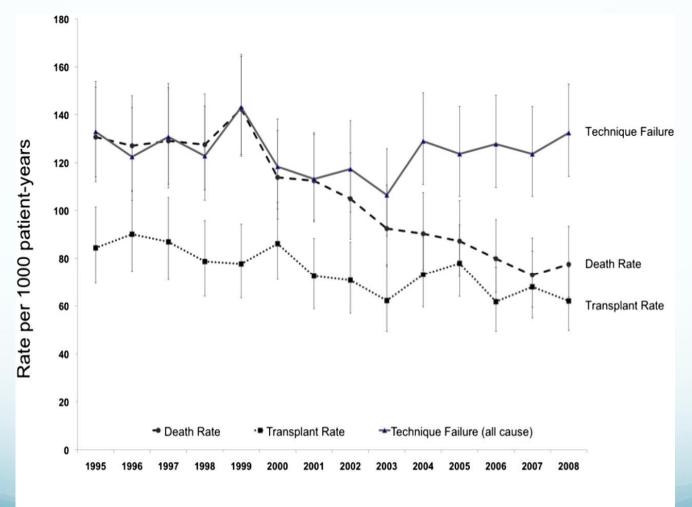
What About Trends In Technique Survival?

Less Patients Need to Transfer in United States



Mehrotra et al, Kidney Int 2009; 76: 97-107

Technique Survival Trends in Canada



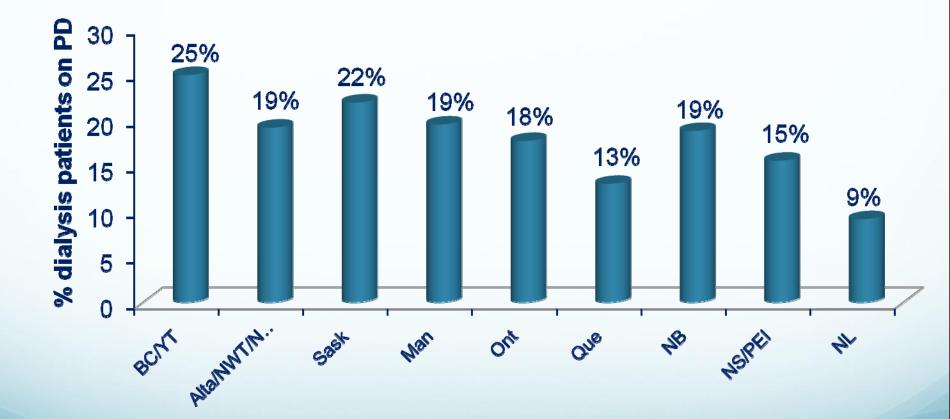
Year of Peritoneal Dialysis Initiation 1995-2008

Perl et al, Clin J Am Soc Nephrol 2012; 7: 1145-54

Geographic Variability in PD Use



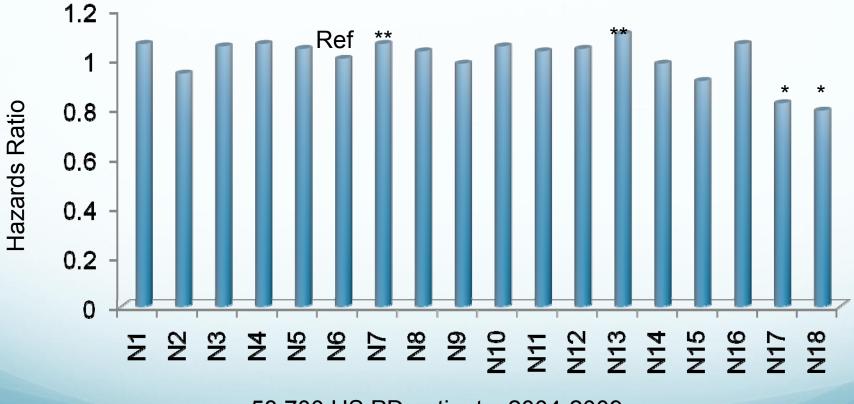
Geographic Variability in PD Use



CORR 2012

Geographic Variability in PD Outcomes

Death/HD Transfer



58,700 US PD patients, 2004-2009

Mehrotra et al, Perit Dial Int 2012; 322-31

Geographic Variability in PD Outcomes

Variables		p-value	HR	CI 95.0	CI 95.0% HR			HR
variables		p-value	пк	Lower	Upper	0	0.5	1 1.5 2
Age > 65 Years		0.19	1.16	0.93	1.43			ı ∔ ●1
Non Caucasian race		0.04	1.26	1.003	1.58			
Educational level	Illiteracy	0.03	1.75	1.04	2.92			\longmapsto
	Elementary	0.02	1.64	1.06	2.54			\longmapsto
	Secondary Higher (reference)	0.05	1.57	0.99	2.49			\longmapsto
Distance from	≤ 25 Km	0.01	1.40	1.07	1.83			
dialysis center	26 - 50 Km	0.36	1.17	0.84	1.63		F	∔ ●───1
	> 50 Km (reference)							
Number of patients	First (<40)	0.03	1.49	1.02	2.18			\longmapsto
per dialysis center	Second (40 - 60)	0.99	0.99	0.68	1.45			4 ──→
(quintiles)	Third (61 - 89)	0.52	1.14	0.77	1.69			↓ ●
	Fourth (90 – 156)	0.24	1.25	0.86	1.81		ŀ	∔ ●1
	Fifth (157-256) (reference)							
Region where patients	South (reference)							
live	Northeast	0.002	1.77	1.23	2.55			
	Southeast	0.29	1.20	0.85	1.68		. F	
	West Center	0.69	1.20	0.47	3.10			
	North	0.34	1.33	0.73	2.41			$ \longrightarrow $
SBP > 140 mm Hg		0.09	0.84	0.68	1.03		ب	н
Dialysis Indication	Patient option (reference)							
	Medical	0.64	1.06	0.83	1.34		F	-↓ → ↓
	Only option	0.08	1.27	0.97	1.66		-	H ⁻ -+1
Diabetes		0.36	0.91	0.74	1.12		E H	•1

Time to Peritonitis

Martin et al, Clin J Am Soc Nephrol, 2011; 6: 1944-51

Geographic Variability

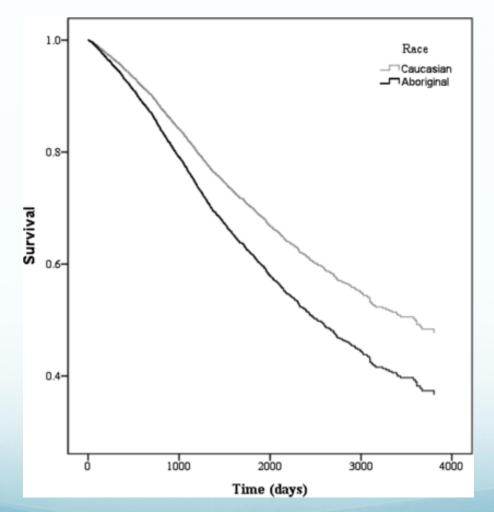
- How well patients do on PD depends on where they live:
 - It is unlikely to be the weather or the health of patients but
 - Either:
 - How PD is practiced and/or
 - Unmeasured differences in access to care
- Need to see what good units do well and what can be improved in units that don't do so well:
 - Need to constantly measure and review outcomes

Race and PD Outcomes United States

- Use of PD, relative to Whites:
 - Significantly lower in Blacks
 - No different in Hispanics

- Blacks and Hispanics:
 - Significantly lower risk for death BUT
 - Higher risk of transfer to hemodialysis (technique failure)

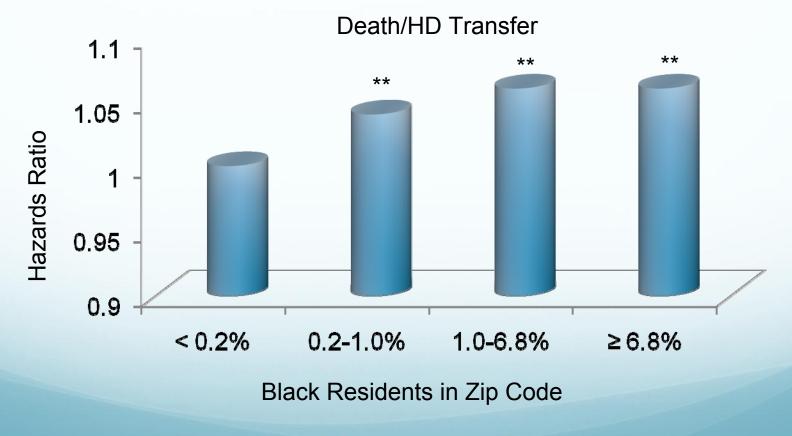
Race and PD Outcomes Canada



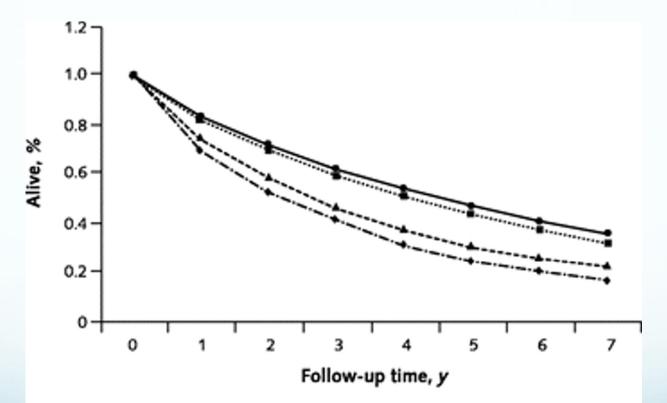
31,576 Canadian PD patients, 2000-'09

Sood et al, Clin J Am Soc Nephrol; 7: 1988-95

Minorities in Neighborhood



Neighborhood Segregation and HD Outcomes



→ Black patients living in ZIP codes with <10% black residents Black patients living in ZIP codes with ≥75% black residents White patients living in ZIP codes with <10% black residents White patients living in ZIP codes with ≥75% black residents

Rodriguez et al, Ann Intern Med 2007; 146: 493-501

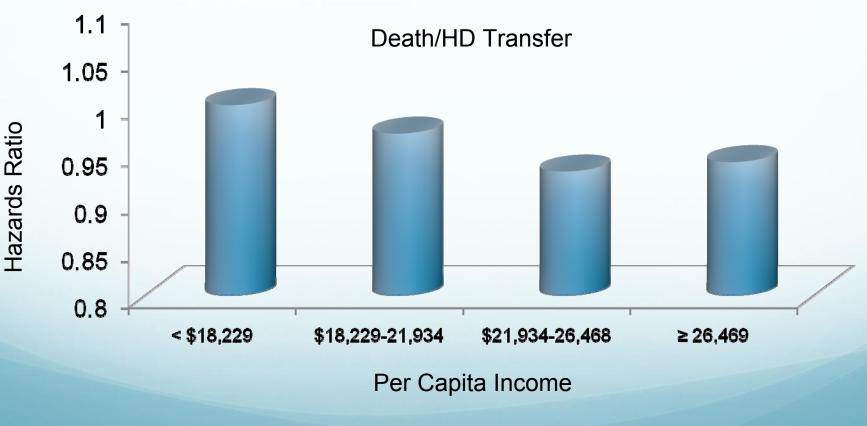
Race and Neighborhood Segregation

- Opportunities exist to improve outcomes of racial/ethnic minorities with PD in US/Canada:
 - Blacks in US: higher technique failure
 - Aboriginals in Canada: higher mortality
- In US, dialysis patients in black-segregated communities don't do so well, either with HD or PD

• Need a different skill set, a different way to engage and educate patients from different racial/cultural backgrounds

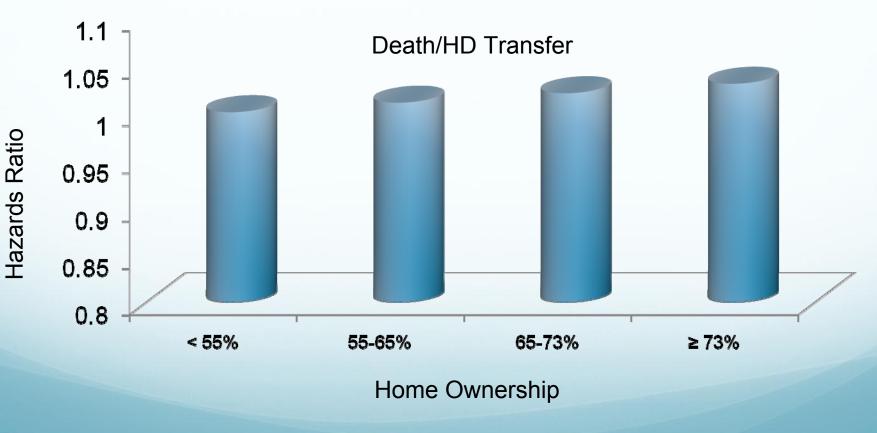
Neighborhood Affluence

No Significant Difference

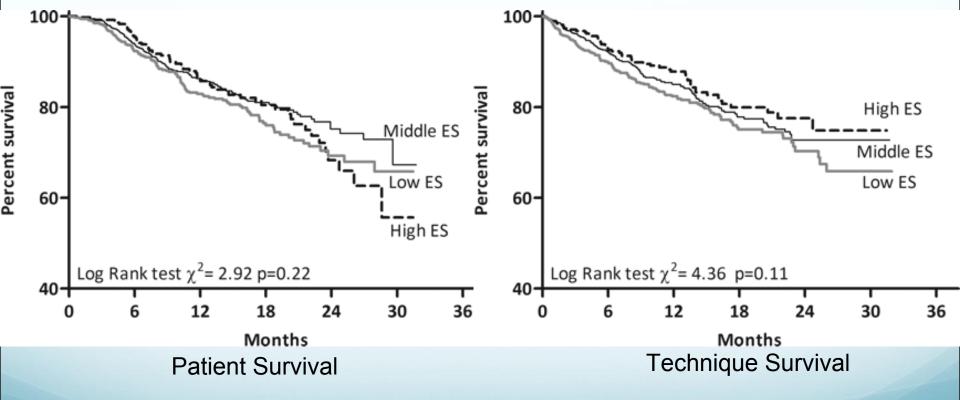


Neighborhood Affluence

No Significant Difference



Family Income and PD Outcomes

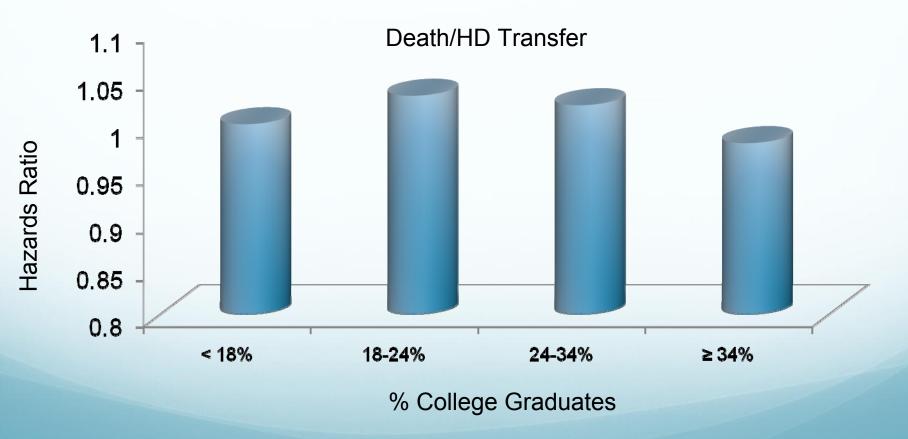


1952 Brazilian PD Patients

Bastos et al, Clin J Am Soc Nephrol 2011; 6: 1676-83

Neighborhood Educational Attainment

No Significant Difference



Educational Attainment and Peritonitis

-	Variables		p-value HR		CI 95.0%				HR	
_	, an above		p .uue		Lower	Upper	0	0.5	1 1.5 2	
	Age > 65 Years		0.19	1.16	0.93	1.43			⊦ ∔∙−-1	
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Γ	Educational level	Illiteracy	0.03	1.75	1.04	2.92			⊢	
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L		Secondary Higher (reference)	0.05	1.57	0.99	2.49			$ \longrightarrow $	
	Distance from	≤ 25 Km	0.01	1.40	1.07	1.83			I I I I I I I I I I I I I I I I I I I	
	dialysis center	26 - 50 Km	0.36	1.17	0.84	1.63			⊢∔ ●───1	
		> 50 Km (reference)								Time to
	Number of patients	First (<40)	0.03	1.49	1.02	2.18			\longmapsto	
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	Dialysis Indication	Patient option (reference)								
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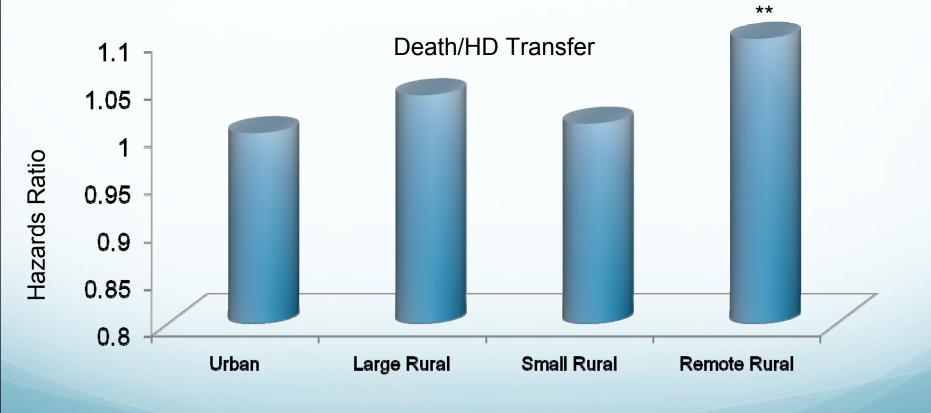
Martin et al, Clin J Am Soc Nephrol, 2011; 6: 1944-51

Economic Status and PD Outcomes

• This seems to be the least important of the sociodemographic factors

- It is possible to achieve good outcomes with PD in individuals that are poor and/or less educated:
 - The major limitation seems to be living conditions

Facility Neighborhood "Rurality"



Rurality/Distance from Facility Canada

	Death	Technique Failure
Rural Residence	1.17 (0.98-1.39)	0.91 (0.74-1.13)
Distance from dialysis unit (every 1 km increase)	1.00 (0.99-1.00)	0.99 (0.99-1.01)

5162 PD patients from Ontario, 1995-2005

Chidambaram et al, Perit Dial Int 2011; 31: 565-73

Distance from Dialysis Facility and Peritonitis Risk

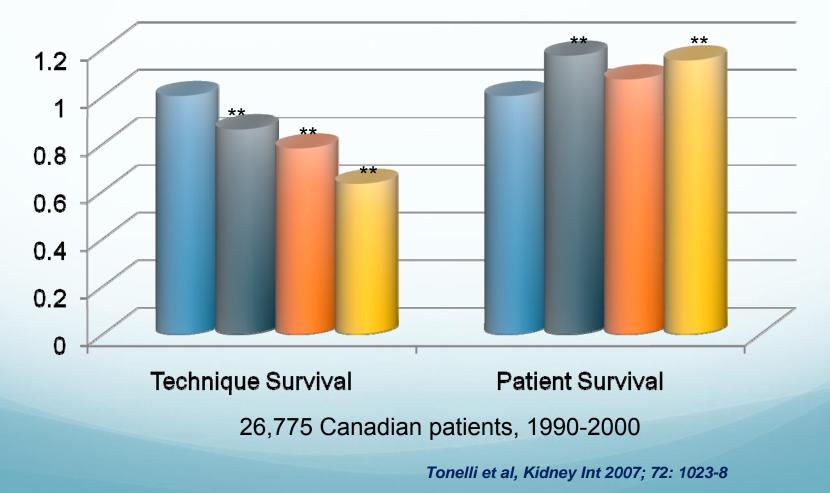
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		0.10		0.03				
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	Secondary	0.05	1.57	0.99	2.49			\longmapsto
	Higher (reference)							
Distance from	≤ 25 Km	0.01	1.40	1.07	1.83			⊢ −•−1
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	North	0.34	1.33	0.73	2.41			• • • • • • • • • • • • • • • • • • •
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-								
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		0.36	0.91	0.74	1.12			Ι.

Time to Peritonitis

Martin et al, Clin J Am Soc Nephrol, 2011; 6: 1944-51

Distance from Unit Canada

■ ≤ 50 km ■ 50.1-150 km = 150.1-300 km = > 300 km



Rurality and Distance from Facility

 Dialysis units in remote rural areas (0.1% of all facilities) seem to face unique challenges that need to be understood

 We need to test interventions, like telemedicine, to improve the outcome of patients that live particularly far away from dialysis facilities

Conclusions

- The dialysis therapy itself does not determine how well patients do, either early or late
- People likely practice PD differently in different parts of the country/world; it is important to identify proven "best demonstrated practices"
- Training and management in PD should be culturally appropriate to overcome unique problems seen in minorities or segregated neighborhoods
- Patient's affluence or educational attainment do not determine how well people do on PD
- Finally, there is a compelling need to identify practices that will allow programs to better support patients who live in remote locations, far from dialysis facilities