WHY ISN'T PD THE FIRST CHOICE FOR YOUR PATIENTS ?

BC NEPHROLOGY DAYS VANCOUVER OCTOBER 2007

Peter G Blake

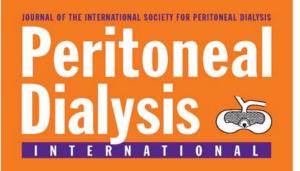
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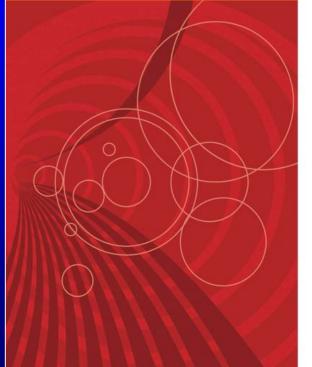
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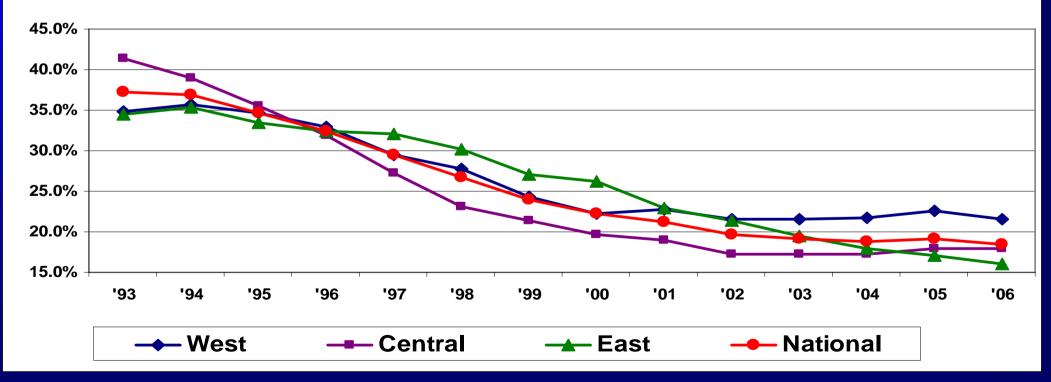
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TOPICS TO COVER

- What proportion of patients should do PD ?
- Which patients should do PD?
- What are the obstacles to getting patients to do PD ?
- How do we grow PD ?

FALL IN PD UTILIZATION CANADA 1993 - 2006

PD Penetraion National, Regional



CAUSES OF DECLINE IN PD USE

- Proliferation of HD units
- Increased patient age and co-morbidity with inability or reluctance to do PD
- Physician concerns about efficacy of therapy

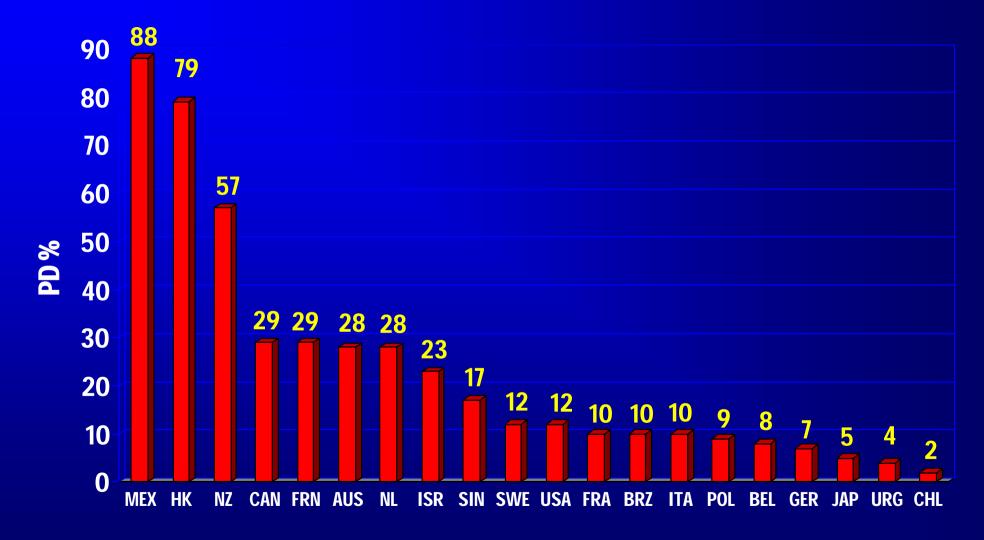
CAUSES OF DECLINE IN PD USE OTHER POSSIBILITIES

- U.S. studies suggesting higher mortality on PD
- Poor training of fellows in PD ?
- PD catheter insertion problems

WHAT PROPORTION OF PATIENTS SHOULD DO PD ?

- Huge variation internationally and nationally
- Driven largely by non-medical factors
- Ontario PD Initiative targets 30% PD
- Any evidence ?

PD Share Distribution Around the Globe



USRDS 1999

Baxter¹99

WHAT PROPORTION OF PATIENTS SHOULD DO PD ?

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- Ontario PD Initiative targets 30% PD
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WHAT PROPORTION OF PATIENTS SHOULD DO PD ?

- Survival data
- Economic perspective
- Patient views
- Nephrologist views

SURVIVAL ON PD VERSUS HD

Has been controversial for years

No randomized trials

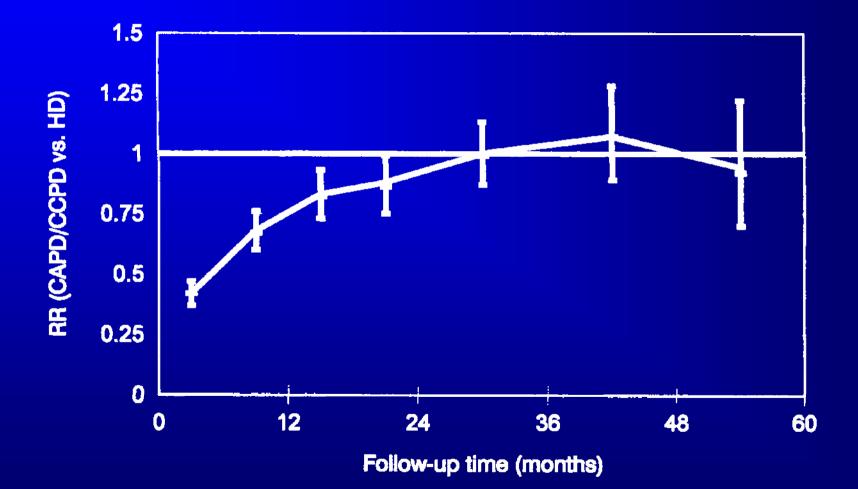
Results depend on methodology

• Results in U.S. tend to differ from those elsewhere

CANADIAN SURVIVAL DATA Fenton et al (PDI 1998) Relative risks for PD vs HD

	AT	ITT
All	0.73*	0.93*
Non DM < 65	0.53*	0.84
DM > 65	0.75*	0.95
Non DM 65 +	0.76*	0.90
DM 65+	0.88	1.04
*95% Cl < 1.0		

Mortality Rate Ratios for PD Relative to HD



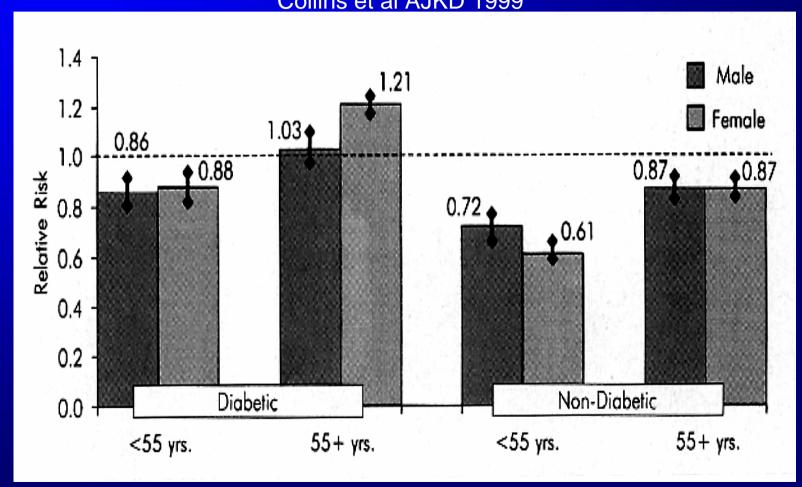
Schaubel et al 1998

U.S. DATA - COLLINS HCFA STUDY AJKD 1999

 ITT based on modality at 90 days with censoring 60 days post switches and 2 year follow up

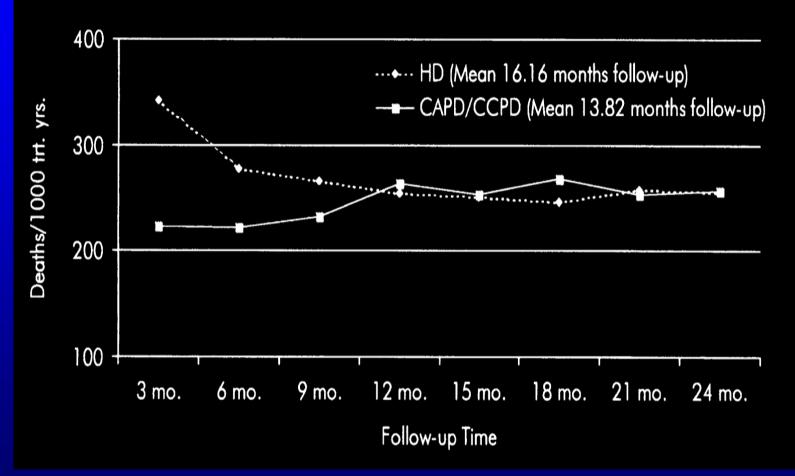
 106,000 incident patients 1994-1998, correction for age and race, diabetics and non diabetic separately

Relative Mortality Risk PD vs HD All Patients by Age, Sex and Diabetes Collins et al AJKD 1999



Unadjusted Mortality Rates PD and HD

Collins et al AJKD 1999



COMPARATIVE SURVIVAL PD AND HD

 More recently, Ganesh et al (JASN 2003) and Stack et al (KI 2003) used incident data from USRDS to show that patients with CHF and with CAD had worse survival on PD, especially if diabetic

RELATIVE RISK OF MORTALITY (PD/HD) IN INCIDENT ESRD WITH CORONARY DISEASE

	RR (unadj)	RR (adj)
DM	1.07	1.23
	NS	P < 0.001
Non DM	1.01	1.20
	NS	P < 0.001

Ganesh JASN 2003

RELATIVE RISK OF MORTALITY (PD/HD) IN INCIDENT ESRD WITHOUT CORONARY DISEASE

	RR (unadj)	RR (adj)	
DM	0.92	1.17	
	P < 0.05	P < 0.001	
Non DM	0.69	0.99	
	P < 0.001	NS	

Ganesh JASN 2003

COMPARATIVE SURVIVAL HD v PD (Vonesh et al KI 2004)

- USRDS 1995-2000, N = 400,000, incident study
- Stratification by age, diabetes and co-morbidity is required
- 55% had comorbidity, 45% had DM and median age was 65
- RR of mortality expressed as HD:PD

HD v PD by Age, DM and Co-morbidity Vonesh et al KI 2004

	No co-morbidity	Co-morbidity
	RR	RR
No DM 18-44	1.24 (1.07-1.44)	1.19 (0.94-1.50)
45-64	1.13 (1.02-1.25)	1.01 (0.92-1.11)
65+	1.13 (1.05-1.21)	0.96 (0.91-1.01)
DM 18-44	1.22 (1.05-1.42)	1.10 (0.92-1.32)
45-64	0.92 (0.85-1.00)	0.82 (0.77-0.87)
65+	0.86 (0.79-0.93)	0.80 (0.76-0.85)24

CHOICE STUDY Jaar et al Ann Intern Med 2005

- National prospective cohort study in U.S.
- NIH/NIDDK funded
- 1041 incident dialysis patients 1995-98
- 81 dialysis centers with oversampling of PD to allow statistical comparison
- Mean follow up 2.4 years (up to 7 years)

THE FIRST PROSPECTIVE COHORT COMPARATIVE HD PD STUDY KI 2000 57: 1720-26

Kidney International, Vol. 57 (2000), pp. 1720-1726

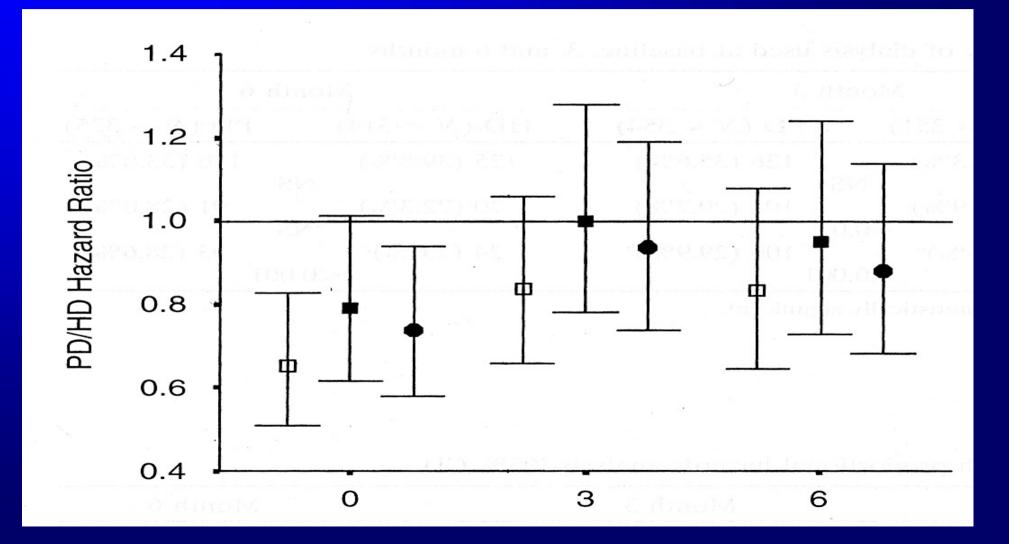
Comparative mortality of hemodialysis and peritoneal dialysis in Canada

SEAN W. MURPHY, ROBERT N. FOLEY, BRENDAN J. BARRETT, GLORIA M. KENT, JANET MORGAN, PAUL BARRÉ, PATRICIA CAMPBELL, ADRIAN FINE, MARC B. GOLDSTEIN, S. PAUL HANDA, KAILASH K. JINDAL, ADEERA LEVIN, HENRY MANDIN, NORMAN MUIRHEAD, ROBERT M.A. RICHARDSON, and PATRICK S. PARFREY

THE FIRST PROSPECTIVE COHORT COMPARATIVE HD PD STUDY Murphy et al (KI 2000 57: 1720-26)

- 822 consecutive incident patients at 11 Canadian centres 1993-1994
- Extensive assessment of status and comorbidity prior to starting dialysis
- Adjustment for demographics and co-morbidity score based on presence and severity of comorbid conditions
- Mean follow up 24 months (up to 56 months)

PD/HD HAZARD RATIOS FOR MORTALITY UNADJUSTED AND ADJUSTED BY COMORBIDITY AT 0, 3 AND 6 MONTHS (Murphy KI 2000)



COMPARISON OF THE TWO PROSPECTIVE COHORT STUDIES

	CHOICE	Murphy et al
Ν	1041	822
No of centres	81	11
Enrolment	10 weeks	0 weeks
Mean follow up	2.4 y	2.0 y
Mean age (yrs)	54 (P), 59 (H)	56 (P), 59 (H)
% PD	26	34, 50, 51
% DM	47	36
% white	66	80 29

PD vs HD DANISH REGISTRY Heaf et al NDT 2002

- Almost 5000 patients treated 1990-1999 35% on PD
- Correction for demographics and recorded comorbidity
- Intent to treat and treatment received analyses
- Survival benefit for PD in first 2 years no difference subsequently

PD versus HD (Heaf et al NDT 2002)

	Intent to treat	As treated
All	0.86 (.5774)	0.65 (.5972)
Non-DM	0.84 (.7594)	0.61 (.5470)
DM	0.93 (.76-1.14)	0.69 (.5785)
Age > 55	0.85 (.7694)	0.66 (.5874)
DM > 55	1.04 (.75-1.43)	0.75 (.5799)

SURVIVAL STUDIES

- HD and PD may have modest survival advantages in particular subgroups but these are unproven
- Advantages, if any, are small relative to larger issues
- Broadly speaking, the 2 modalities have equal survival overall

COSTING STUDIES

- Every comparative study in the developed world shows that PD costs substantially less than HD
- The difference in Canada is c 40%
- Cost is not the only factor and we treat individual patients but if all else is equal.....

COMPARATIVE DIALYSIS COSTS BY MODALITY IN CANADA (Goeree et al 1995)

	Centre HD	Self-care HD	Home HD	CAPD
Salaries	\$19,676	\$16,100	\$6,780	\$6,670
Supplies	\$10,779	\$8,236	\$9,013	\$13,836
Medicines	\$6,226	\$4,032	\$2,115	\$3,547
Others	\$24,474	\$7,865	\$8,141	\$7,865
Total	\$54,929	\$43,313	\$26,048	\$31,918

PATIENT VIEWS

- Prichard et al (PDI 1996) 150 new starts in Montreal
- 31 directed to HD and 14 to PD
- 31 diabetics encouraged to do PD 17 did
- The remaining 74 could do either and got free choice split was 50:50

NEPHROLOGIST VIEWS

- Jung et al (PDI 1999) surveyed all CSN members – 66% replied and recommended HD PD Split was 66:34
- Mendelssohn et al (AJKD 2001) surveyed US nephrologists and recommended split was 71:29
- Jassal et al (NDT 2002) studied UK nephrologists and suggested split was 62:38

WHAT PROPORTION SHOULD DO PD?

- Survival is broadly similar
- PD costs less
- Half of patients choose PD if given informed choice ?
- Nephrologists think about one third should do PD
- So what is the problem !?

WHAT PROPORTION OF PATIENTS CAN DO PD ?

- Oliver et al (KI 2007) studied 134
 incident patients in Toronto
- 81% had at least 1 potential PD barrier
- With home care support 80% could do PD versus 65% without

Oliver et al (KI 2007)

- Of eligible patients, in areas with home care 59% chose PD versus 58% in areas without home care
- However, the actual utilization of PD was 47% in the home care areas versus 37% in areas without home care

WHICH PATIENTS ?

- About 20-30% have a strong contraindication
- Another 20-30% have significant barriers that can be addressed
- The rest are good PD candidates

MAJOR CONTRAINDICATIONS

- Previous major abdominal surgeries
- Ostomies
- Morbid obesity
- Unable to do procedures and no one else to help
- Unwilling to do it

RELATIVE CONTRAINDICATIONS

- Major medical co-morbidity
- Psychiatric illness
- Impaired vision or dexterity
- Poor living conditions
- Unsupportive relatives

• So, what is the problem ?

• Why is PD so hard to grow ?

OBSTACLES TO PD GROWTH

- Late referrals and 'parachutes'
- Poorly structured pre dialysis care and education
- Patient procrastination and denial re modality education and selection
- Patient fear and lack of confidence

OBSTACLES TO PD GROWTH

- Physician biases
- Patient body image issues
- Long wait lists for catheters
- High catheter failure rates
- High technique failure

SOLUTIONS TO BARRIERS

- Well organized pre dialysis education
- Insistence on patient and family participation
- Tours of PD unit and meetings with PD nurses and patients for all suitable patients
- Avoid biases or 'damning by faint praise' !

SOLUTIONS TO BARRIERS

- Constant encouragement to do PD neutral approach is not enough !
- Follow up post PD selection until dialysis needed
- Apply modality education process to all late referrals, parachutes etc
- Use information technology to do ongoing CQI on the modality selection process

SOLUTIONS TO BARRIERS

PD catheters

PD patient retention

PD CATHETERS



PD CATHETERS

- Need for a 'champion' physician, surgeon or radiologist
- Ideal person and technique will vary with centre

PD PATIENT RETENTION

- Peritonitis CQI initiative needed
- Catheters likewise
- Patient and Family 'Burnout'



ONTARIO PD INITIATIVE

- Initiative by Ministry of Health to reverse decline in PD
- Headed by Dimitrios Oreopoulos and Sandra Coleman
- See PDI Sept/Oct 2007
- Article by Oreopoulos and Coleman and Commentary by Jindal

ONTARIO PD INITIATIVE RECOMMENDATIONS

- Increase PD use to 30 % over 2-3 years – target based on physician and patient surveys and cost issues
- For home HD centers total home dialysis should be 40%

ONTARIO PD INITIATIVE COMPONENTS

- Early referral
- Pre dialysis education
- Prompt expert catheter placement
- Data management

ONTARIO PD INITIATIVE IMPROVE PD RETENTION

- CCAC home visits to support PD
- Nursing homes able to deliver PD one per regional kidney centre

ONTARIO PD INITIATIVE THE RESPONSE

- Variable response from nephrologists
 and renal programs
- Concerns re unrealistic targets and time frames
- Concerns re penalties for centres not meeting targets
- Concerns re costs

PD PRESCRIPTION

- If we want to keep patients on PD we need to prescribe it intelligently
- Previous high Kt/V approach was often not lifestyle friendly built is no longer required
- Aiming for Kt/V 1.7 per week is not difficulty

PD PRESCRIPTION

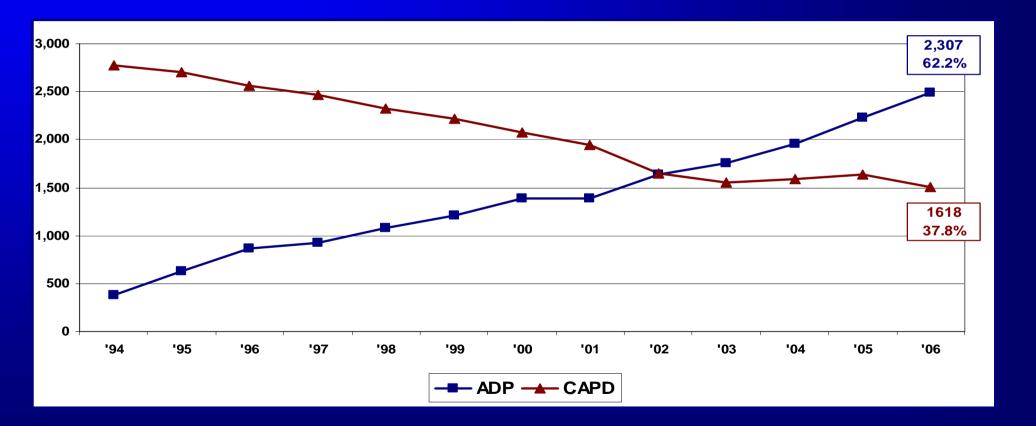
- However, we need to pay attention to volume control in patients
- We also need to be aware of the toxicity of hypertonic glucose
- In particular, think of patient and caregiver lifestyle



APD



SWITCH FROM CAPD TO APD CANADA 1994 - 2006



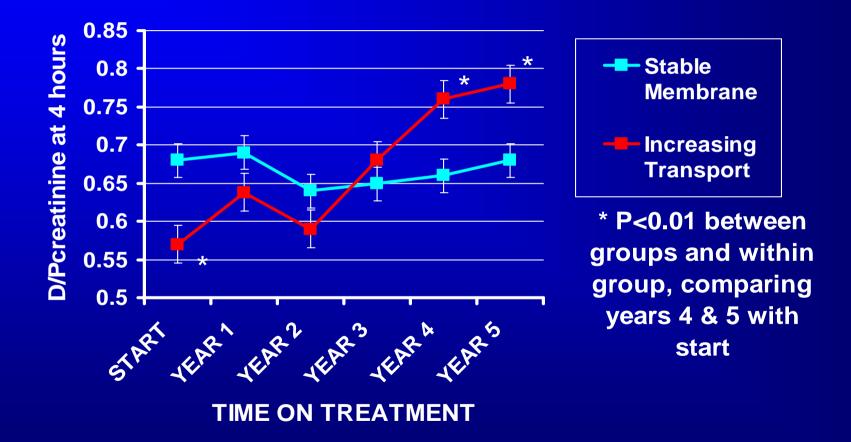
SWITCH FROM CAPD TO APD

- Mainly driven by convenience and lifestyle factors – for patients and caregivers
- To a lesser extent for medical reasons
 clearance and volume reasons or for high transporters
- Cost issues are also a factor

GLUCOSE SPARING STRATEGIES

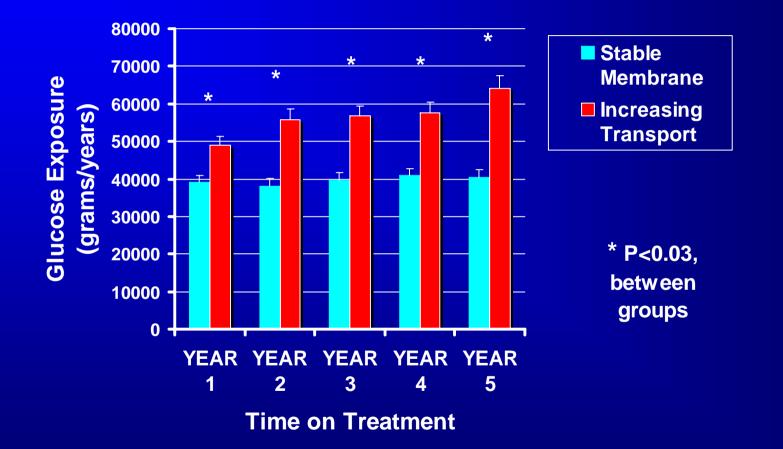
- Increasing body of evidence that hypertonic glucose is toxic to the peritoneal membrane and may lead to type I membrane failure
- Concerns about effects of glucose absorption on the cardiovascular risk profile

Longitudinal Membrane Transport



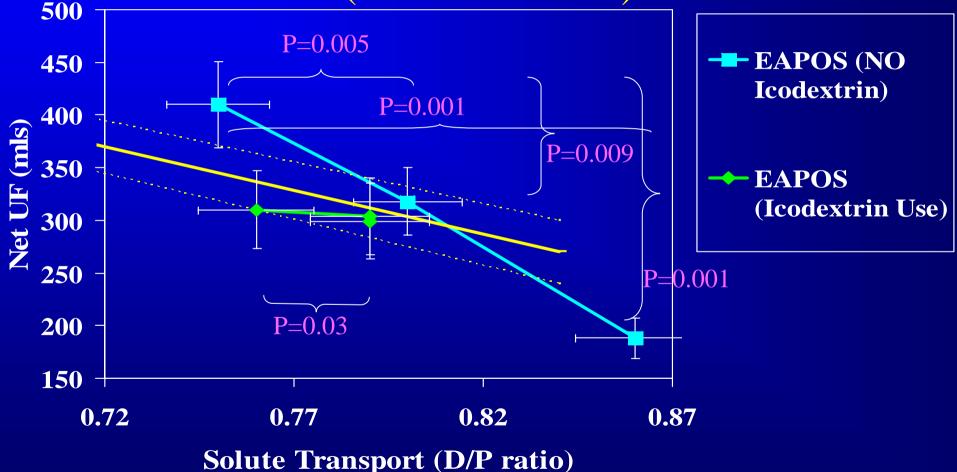
Davies et al, JASN 12:1046-51, 2001

Yearly Peritoneal Glucose Exposure



Davies et al, JASN 12:1046-51, 2001

EAPOS: Influence of Icodextrin on evolution of solute transport and UF capacity (paired data) (Davies KI 2005)



SYSTEMIC EFFECTS OF GLUCOSE ABSORPTION



SYSTEMIC TOXICITY OF GLUCOSE IN PD PATIENTS

- Exacerbation or induction of diabetes
- Hyperinsulinemia
- Promotion of obesity
- Decreased appetite
- Increases in hyperlipidemia (LDL and TGs)
- Increase in cardiovascular risk

GLUCOSE SPARING STRATEGIES THE PARADOX

- These are based on minimizing hypertonic glucose exposure
- But volume control often requires
 greater hypertonic glucose exposure
- There is an apparent contradiction here needing resolution

GLUCOSE SPARING STRATEGIES

- APD can be viewed as a glucose sparing strategy – less glucose exposure per volume of fluid removed
- Icodextrin and i/p amino acids are also glucose sparing
- Salt and water restriction decrease the need for hypertonic glucose

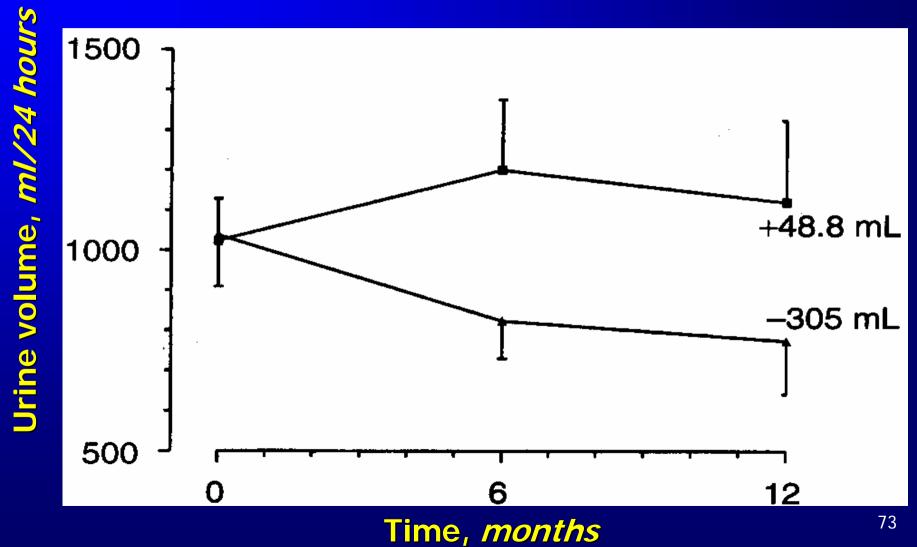
GLUCOSE SPARING STRATEGIES

- Strategies that preserve residual renal function decrease need for hypertonic glucose
- A number of these have become apparent from recent studies

Medcalf et al KI 2001

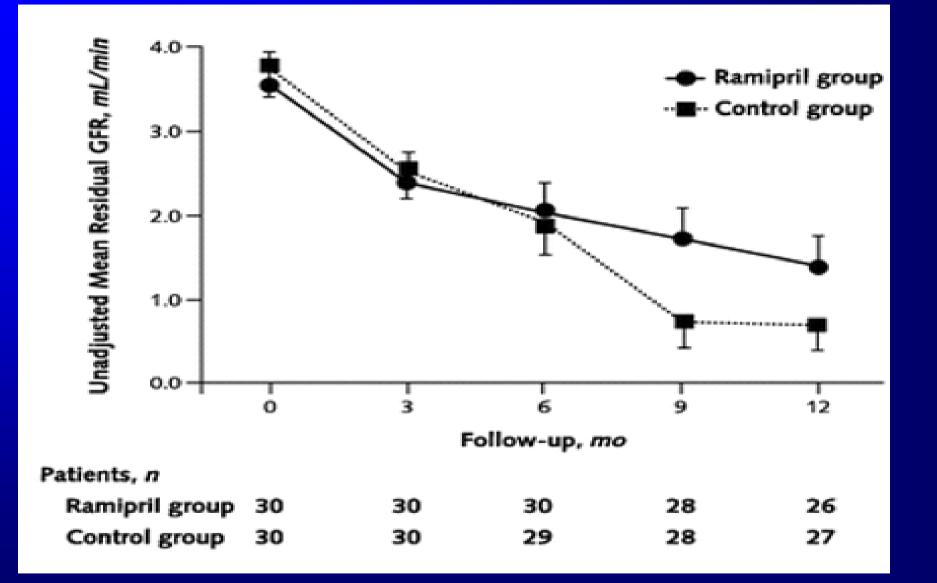
- Open label RCT of 61 incident PD patients at a single centre randomized to Furosemide 250 od or to control group
- Urine volume maintained in F group but fell in controls (p< .05)
- Urine Na excretion greater in F group (p.04)
- % body water stable in F group but rose in controls (52 vs 64%, p 0.1)

Effect of Furosemide on urine output in CAPD (Medcalf et al KI 2001)



Li et al (Ann Int Med 2003)

- Single centre, open label RCT
- 60 prevalent CAPD patients with GFR > 2 ml/min, BP > 120/70, no ACEI or ARB x 6 mths, no CHF etc (72/217 eligible)
- Ramipril 5 od in treatment group, same 135/85 BP target in each group - 12 month follow up
- Repeated measures analysis of covariance



GLUCOSE SPARING STRATEGIES COMMON CLNICAL ERRORS

- It is important to revise target weight up if patients gain body weight on PD
- Otherwise patient may use hypertonic glucose to try and remove body fat
- This may cause volume depletion and promote further obesity

GLUCOSE SPARING STRATEGIES COMMON CLNICAL ERRORS

- All shortness of breath on exertion is not pulmonary edema Is it de-conditioning ?
- All ankle swelling is not fluid overload.
 Is it new ? Is there another reason ?

GLUCOSE SPARING STRATEGIES COMMON CLNICAL ERRORS

- It is easy to reduce target weight by writing an order on a chart
- You must tell the patient how to do it more hypertonics or salt and water restriction or diuretics...?
- Many patients think you are asking them to lose body weight by eating less

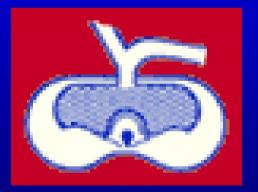
GLUCOSE SPARING vs VOLUME CONTROL

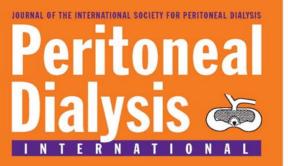
- So, strategies that minimize salt and water intake, preserve residual renal function and make use of icodextrin plus APD are consistent with both glucose sparing and volume control
- However, hypertonic glucose is still often indicated

CONCLUSION

- Growing PD makes sense but is hard to do
- Needs a multifaceted, multidisciplinary approach
- Need to prescribe PD intelligently thinking of clearance, volume and hypertonic glucose minimization and lifestyle

ISPD







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