

How to evaluate the peritoneal membrane?



B. Bammens

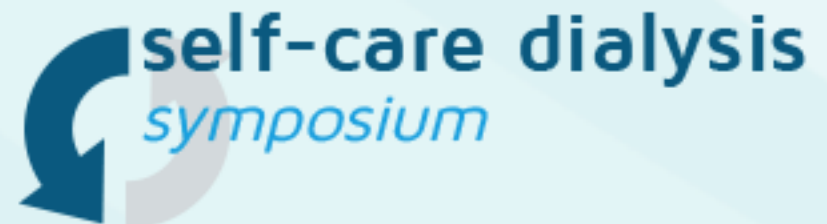
Brussels, May 12 2016

UZ
LEUVEN

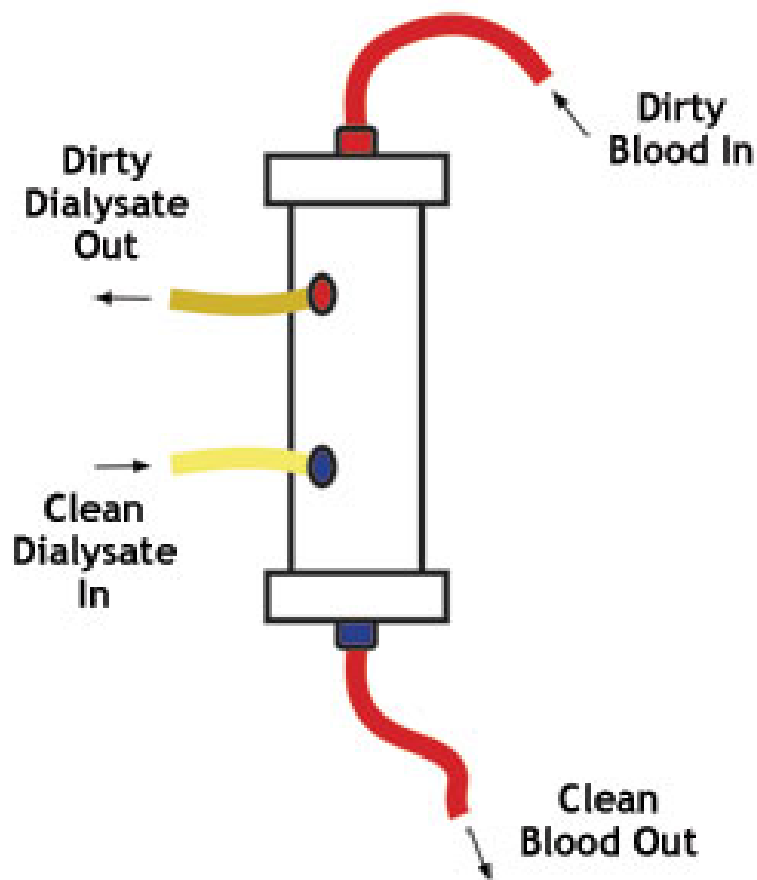
KU LEUVEN

BELGIUM

3rd self-care
dialysis symposium
12th & 13th May 2016

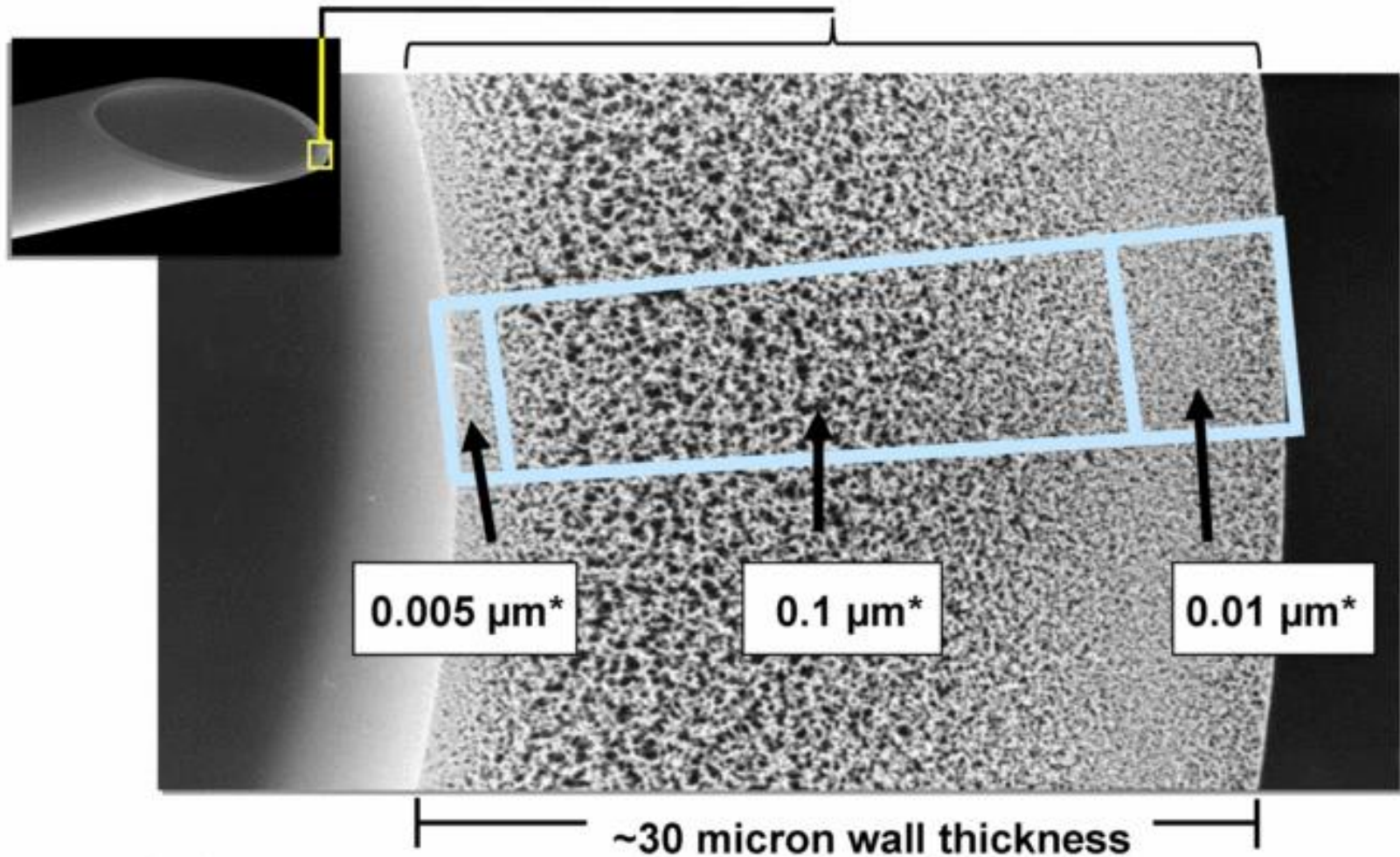


How to evaluate a hemodialyzer?





How to evaluate a hemodialyzer?



* pore size in microns (μm)



How to evaluate a hemodialyzer?

DEFINITIONS OF FLUX, PERMEABILITY, AND EFFICIENCY

Flux

Measure of ultrafiltration capacity

Low and high flux are based on the ultrafiltration coefficient (K_{uf})

Low flux: $K_{uf} < 10$ mL/h/mm Hg

High flux: $K_{uf} > 20$ mL/h/mm Hg

Permeability

Measure of the clearance of the middle molecular weight molecule (eg, β_2 -microglobulin)

General correlation between flux and permeability

Low permeability: β_2 -microglobulin clearance < 10 mL/min

High permeability: β_2 -microglobulin clearance > 20 mL/min

Efficiency

Measure of urea clearance

Low and high efficiency are based on the urea K_0A value

Low efficiency: $K_0A < 500$ mL/min

High efficiency: $K_0A > 600$ mL/min

K_0 —mass transfer coefficient; A —surface area.



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SOLUTE REMOVAL

pore size

surface area



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SOLUTE REMOVAL

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ULTRAFILTRATION

Hemodialyzer vs. peritoneum?

Performance Data

Sieving coefficients of FX Cordiax High-Flux Dialyzers and Hemofilters	Molecular weight (Dalton)	
Albumin	66,500	< 0.001
Myoglobin	17,053	0.5
β_2 -microglobulin	11,731	0.9
Inulin	5,200	1
Membrane material		Holocene® plus
Sterilisation method		INLINE steam
Housing material		Polypropylene

Product Code:	REVACLEAR	REVACLEAR 300	REVACLEAR MAX	REVACLEAR 400
Product Code:	110633	114745	110634	114746
Hemodialysis <i>Q_B = 500 mL/min, UF=0 mL/min</i>				
Q _B (mL/min)	200 300 400 500	200 300 400 500	200 300 400 500 600	200 300 400 500 600
Urea	196 271 321 353	196 272 323 356	198 282 339 376 400	198 281 338 375 401
Creatinine	189 250 289 316	191 256 298 326	195 265 311 341 362	195 267 315 348 370
Phosphate	185 239 274 298	185 242 278 303	191 256 297 324 343	191 255 297 326 346
Vitamin B ₁₂	144 170 186 197	146 174 191 204	158 191 211 225 235	158 191 213 228 240
Hemodialysis <i>Q_B = 400 mL/min, UF=0 mL/min</i>				
Q _B (mL/min)	200 300 400 500	200 300 400 500	200 300 400 500 600	200 300 400 500 600
Urea	199 286 355 408	199 286 355 408	200 293 371 432 479	199 292 369 430 477
Creatinine	194 269 324 364	195 273 330 373	197 281 345 393 430	198 283 348 398 437
Phosphate	191 259 307 343	191 260 309 345	196 273 330 373 406	195 272 330 373 406
Vitamin B ₁₂	154 187 208 223	155 189 212 228	169 211 240 260 276	167 208 236 256 272
Specifications				
Blood flow rate (mL/min)	200-500	200-500	200-600	200-600
Dialysate flow (mL/min)	800	300-800	800	300-800
Membrane				
Material	PAES/PVP	PAES/PVP	PAES/PVP	PAES/PVP
Surface area (m ²)	1.4	1.4	1.8	1.8
UF Coefficient in vitro (mL/h.mmHg) (Bovine blood, hematocrit=32%, protein=60 g/l, 37°C)	50	48	60	54
Priming volume (mL)	84	74	100	93
Residual blood volume (mL)	<1	<1	<1	<1
Fiber Dimensions				
Wall Thickness Membrane (µm)			35	
Inner Diameter Hollow Fiber (µm)			190	
Maximum TMP (mmHg)				
			600	
Sterilization agent				
			Steam	
Sieving Coefficient in vitro. Typical values measured with REVACLEAR Dialyzer according to EN1283				
Vitamin B ₁₂			1.0	
Inulin			1.0	
β_2 -microglobulin			0.7	
Albumin			<0.01	

vs.



+ the effects of
intercurrent disease
time
glucose exposure
...



Peritoneal membrane evaluation

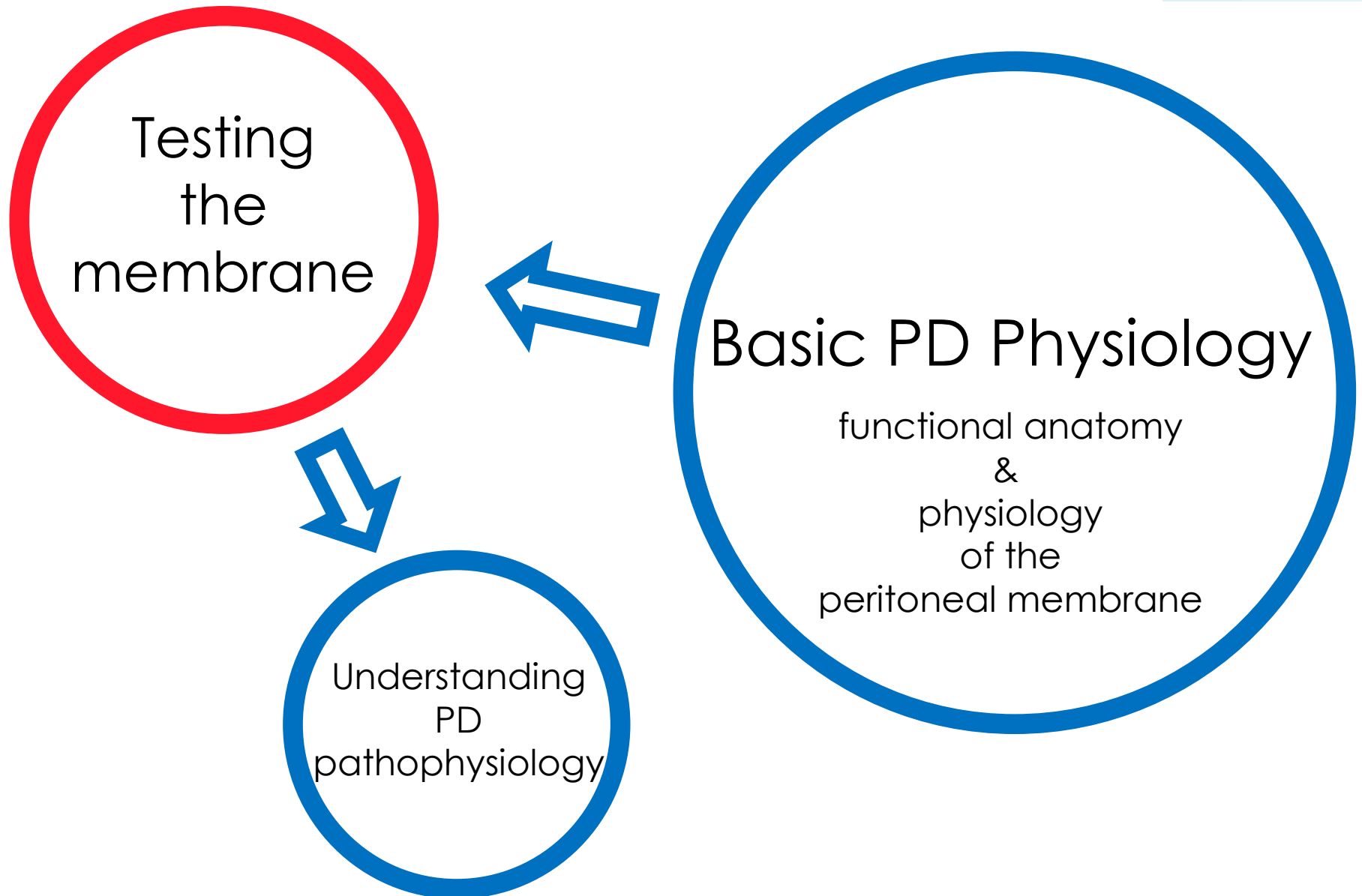


Testing
the
membrane

Testing the membrane

Test	Application/advantages	Limitations
Original PET (2.27%) [12,13,38-40]	Small-solute transport, expressed as D/P value Categories fast/average/slow should guide prescription management (see Table 1) Widely used Definition of UF failure	Limited information No information on sodium sieving, FWT or OC
Modified PET (3.86% glucose) [41]	Small-solute transport, expressed as D/P value Categories fast/average/slow should guide prescription management (see Table 2) Information on sodium sieving Recommended for definition of UF failure	No quantitative information on FWT or OC
APEX (accelerated peritoneal examination test) [42]	Apex time , being the moment when the curves of D/D_0 glucose and D/P_{crac} cross Very suitable to define 'optimal dwell time' for individual patients	No information on sodium sieving, FWT or OC
PDC® (Peritoneal Dialysis Capacity) test [34,35,43-45]	More reliable data because more measuring points Small-solute transport, expressed as area over diffusion distance (A_D/DX). Easily convertible to D/P values Large pore flow Estimate of net peritoneal fluid loss (peritoneal reabsorption) Computer-aided prescription management	Multiple laboratory test needed Computer support for calculations needed
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FWT, free water transport; OC, osmotic conductance.



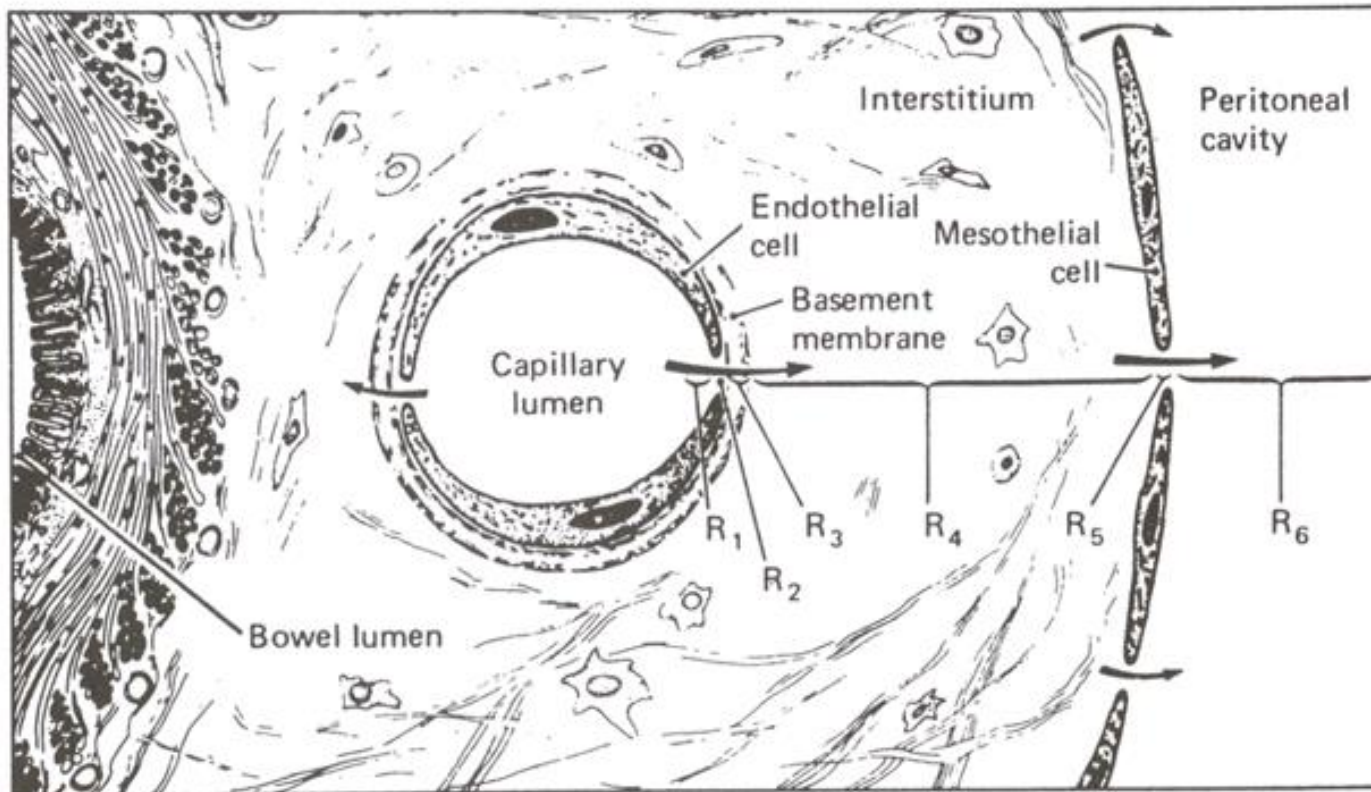
'6 barriers for transport'

Stagnant layers at mesothelial and capillary side: not relevant

Mesothelial cell layer: not relevant

Interstitial tissue: (minor) diffusive resistance

Capillary wall: most important restriction barrier

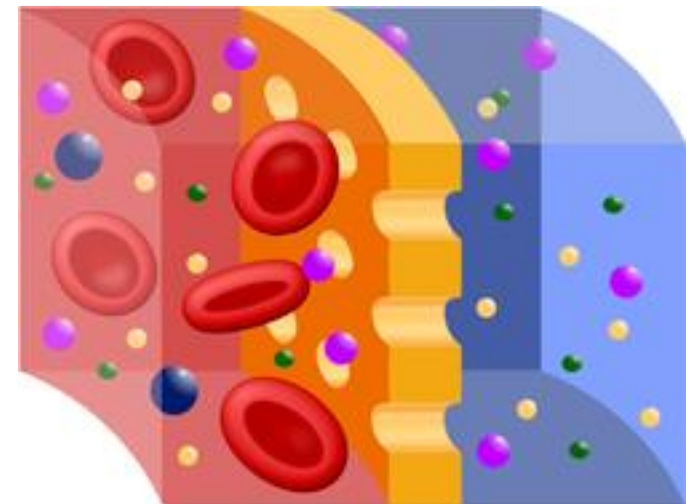
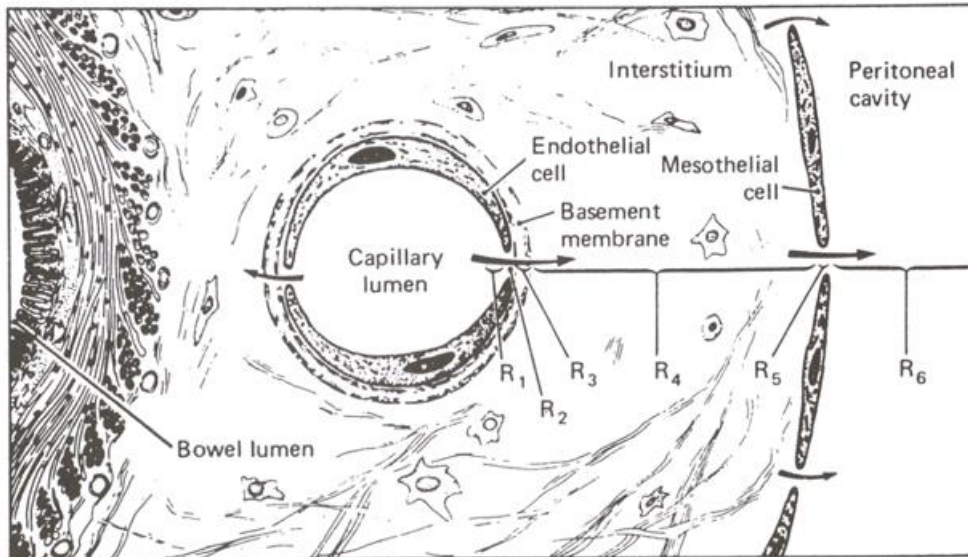




'2D membrane with pores'

Capillary wall is the most important restriction barrier and determines the peritoneal membrane's size-selectivity through a system of pores

→ the "PORE THEORIES"



SOLUTE REMOVAL

ULTRAFILTRATION

pore size

surface area



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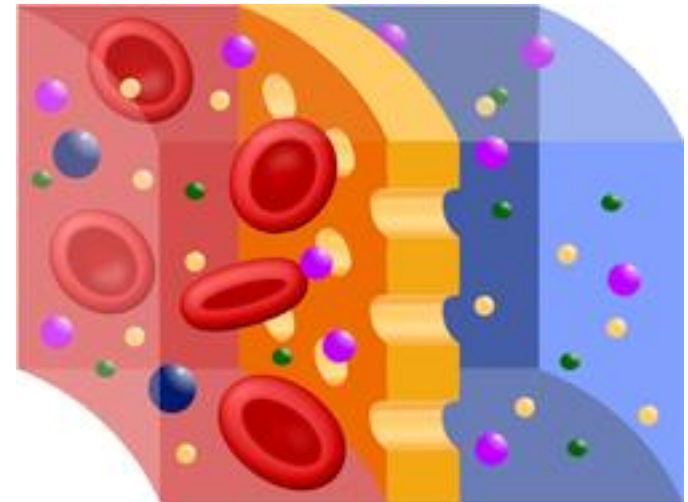
K_o —mass transfer coefficient; A—surface area.

SOLUTE REMOVAL

pore size

surface area

ULTRAFILTRATION



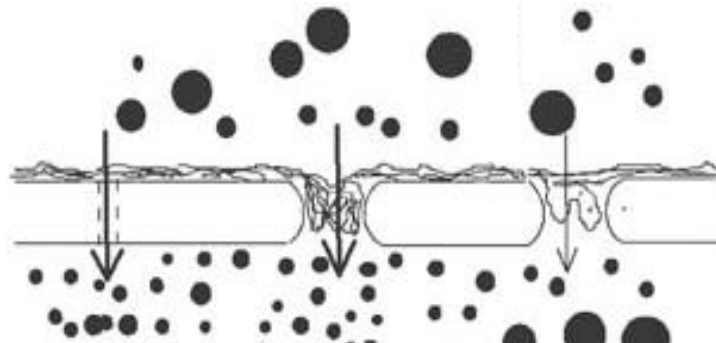


The THREE pore theory

Small pores with constant radius 40-50Å
majority
for transport of low molecular weight solutes

Large pores with various radii, average $> 150\text{\AA}$
minority (less than 0.1% of total pore count)
for transport of macromolecules

Ultra-small pores with radius 3-5Å
for transport of water only
accounts for 1/2 of transcapillary water transport

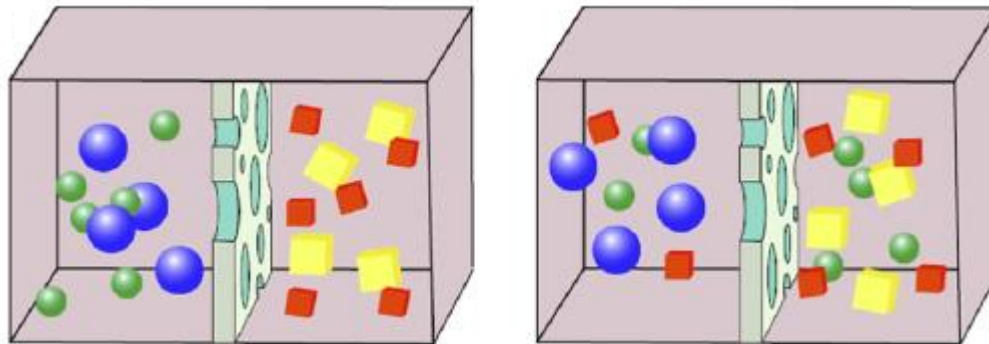


The pore theory explains the “classical” mechanisms of transmembrane transport of molecules.

SOLUTE REMOVAL

DIFFUSION

movement of solutes along their concentration gradient



$$J_s = \frac{D_f}{\Delta x} \cdot A \cdot \Delta C \quad (\text{Fick's first law of diffusion})$$

diffusive permeability (membrane- and solute-specific)

$$J_s = \frac{D_f}{\Delta x} \cdot \mathbf{A} \cdot \Delta C \quad (\text{Fick's first law of diffusion})$$

diffusive permeability (membrane- and solute-specific)

surface area (membrane-specific)

$$J_s = \frac{D_f}{\Delta x} \cdot A \cdot \Delta C \quad (\text{Fick's first law of diffusion})$$

diffusive permeability (membrane- and solute-specific)

surface area (membrane-specific)

concentration difference between plasma and dialysate

$$J_s = \frac{D_f \cdot A \cdot \Delta C}{\Delta x} \quad (\text{Fick's first law of diffusion})$$

diffusive permeability (membrane- and solute-specific)

surface area (membrane-specific)

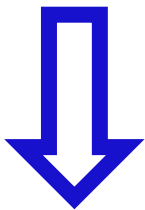
concentration difference between plasma and dialysate

mass transfer area coefficient (MTAC)

$$J_s = \frac{D_f}{\Delta x} \cdot A \cdot \Delta C \quad (\text{Fick's first law of diffusion})$$

$$J_s = \text{MTAC} \cdot \Delta C$$

Transport of small molecules up to MW of $\beta_2\text{M}$ (11,8 kDa)
NOT limited by size of the pores



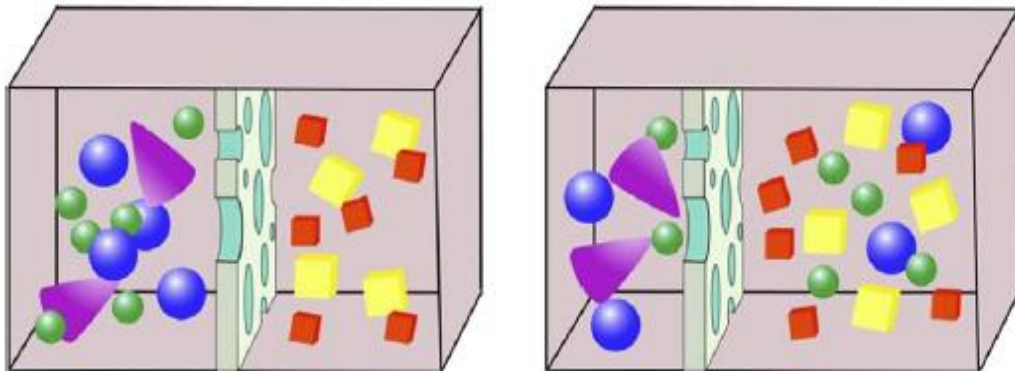
MTAC for a given solute **ONLY** determined by
effective vascular peritoneal surface area (number of pores)

The pore theory explains the “classical” mechanisms of transmembrane transport of molecules.

SOLUTE REMOVAL

CONVECTION

movement of solutes along with fluid as it moves across the membrane (solvent drag)





Convective transport

$$J_s = J_v \cdot \bar{C} \cdot (1 - \sigma)$$

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water flux (membrane-specific)

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mean solute concentration in the membrane $(P+D)/2$



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water flux (membrane-specific)

mean solute concentration in the membrane $(P+D)/2$

Staverman's reflection coefficient
= how difficult it is for a solute to be transported by solvent drag across a semi-permeable membrane
(membrane- and solute-specific)



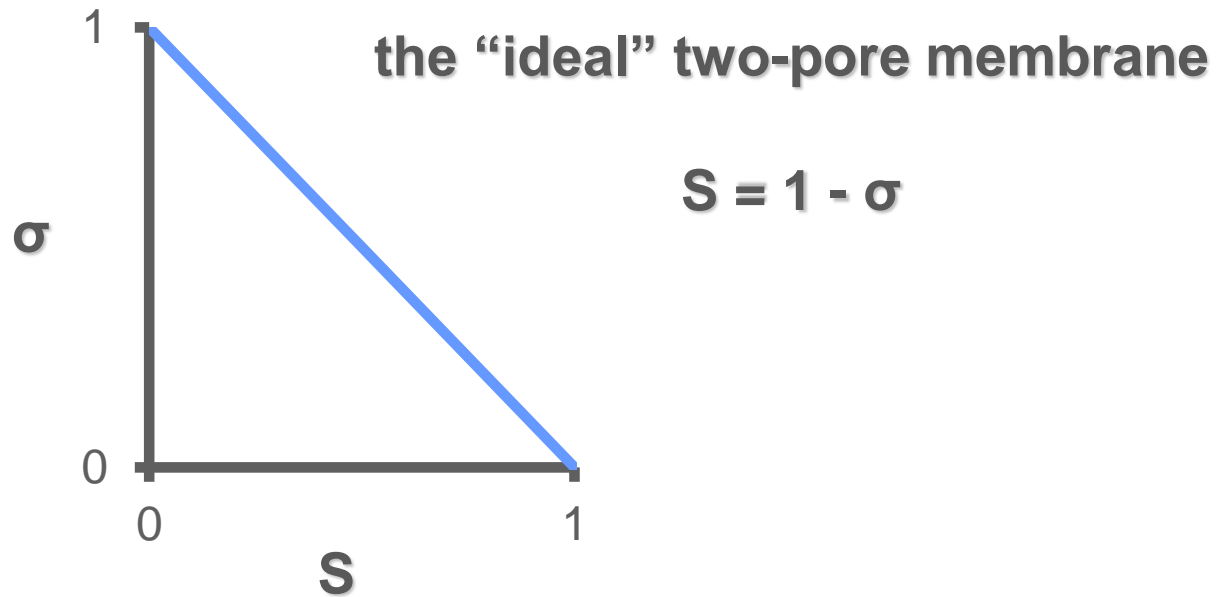
Convective transport

σ **Staverman's reflection coefficient**
= how difficult it is for a solute to be transported by solvent drag across a semi-permeable membrane



S **sieving coefficient**
= how easy it is for a solute to be transported by solvent drag across a semi-permeable membrane

For a semi-permeable membrane, S and σ are expected to be perfectly interchangeable concepts!





Convective transport

σ Staverman's reflection coefficient

= *how difficult it is for a solute to be transported by solvent drag across a semi-permeable membrane*

= **fraction of maximal osmotic pressure a solute can exert across a semi-permeable membrane**

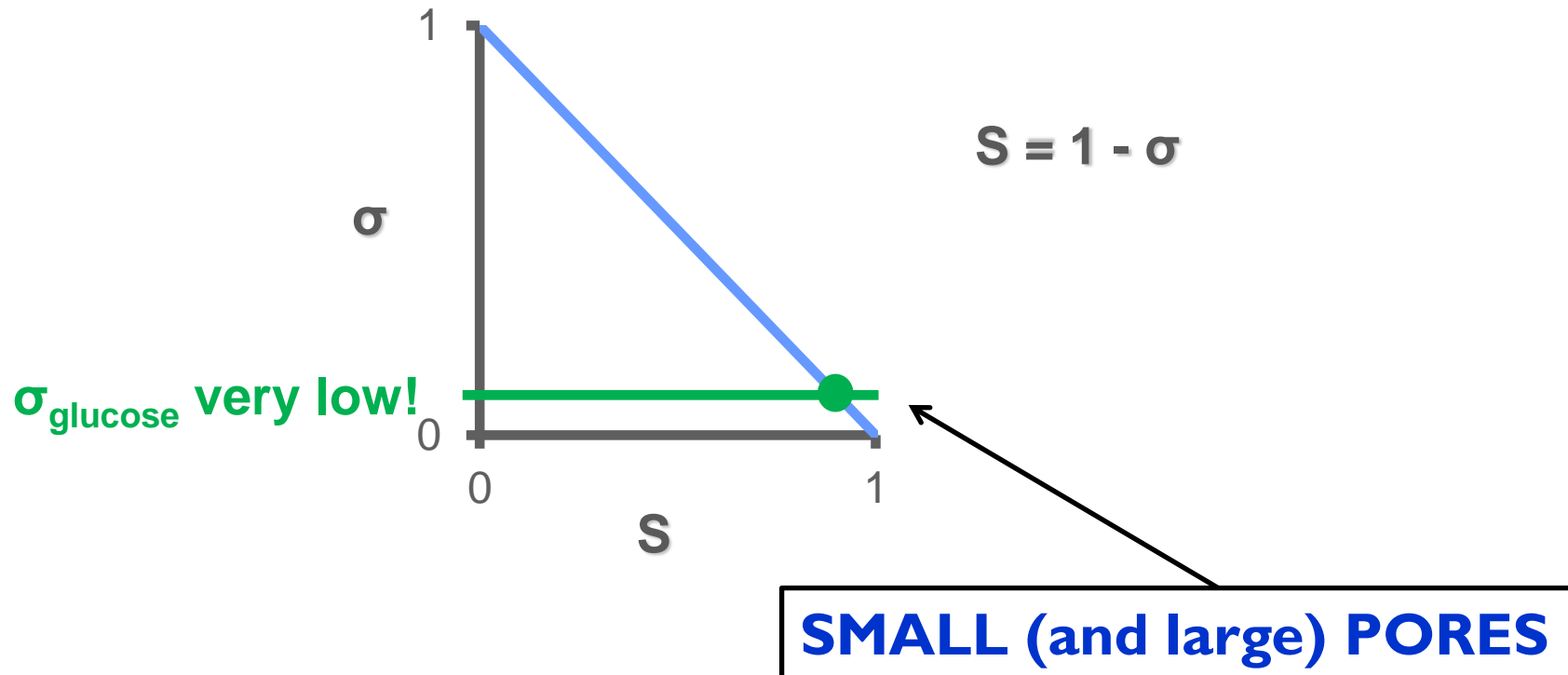


S sieving coefficient

= *how easy it is for a solute to be transported by solvent drag across a semi-permeable membrane*

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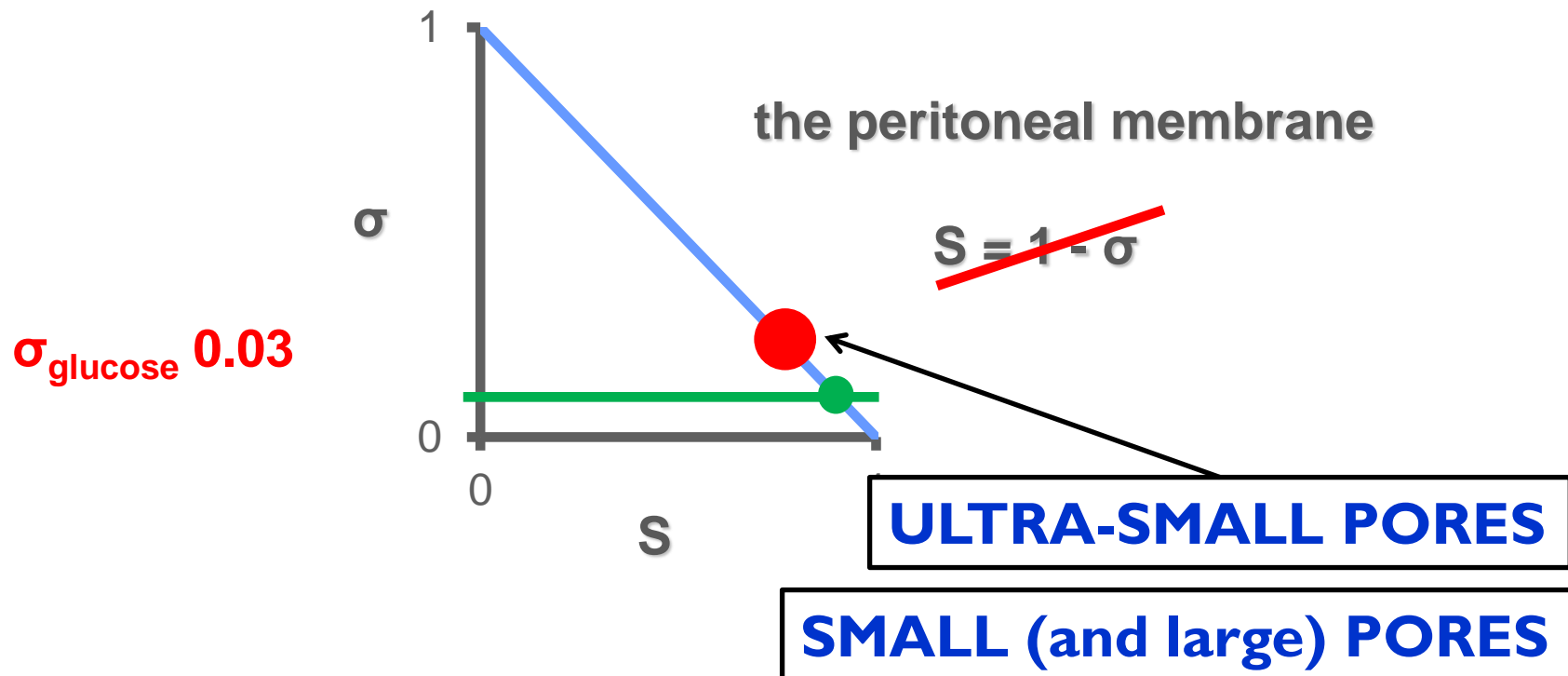
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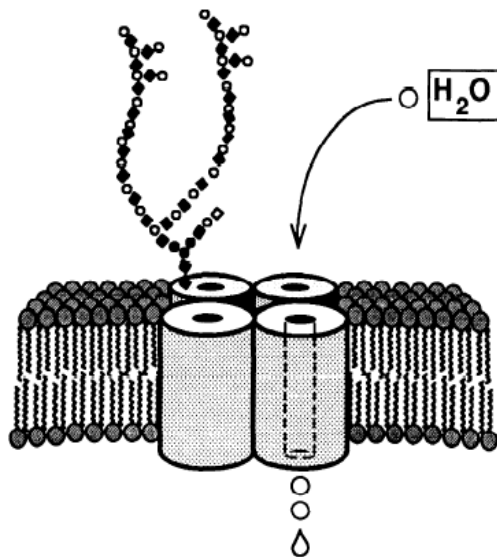
Apparent $\sigma_{\text{glucose}} = \text{higher}$

For a semi-permeable membrane, S and σ are expected to be perfectly interchangeable concepts!

However, the water-only channels make the peritoneal membrane “more than a semi-permeable membrane”!



Ultra-small pores with radius 3-5Å



AQUAPORIN-1

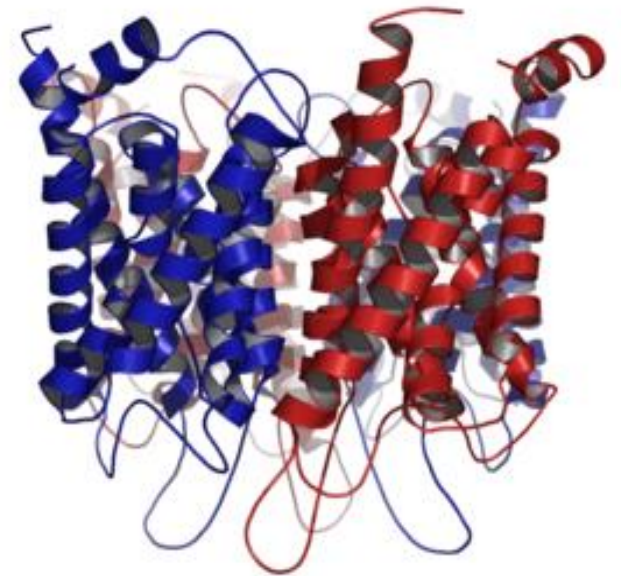
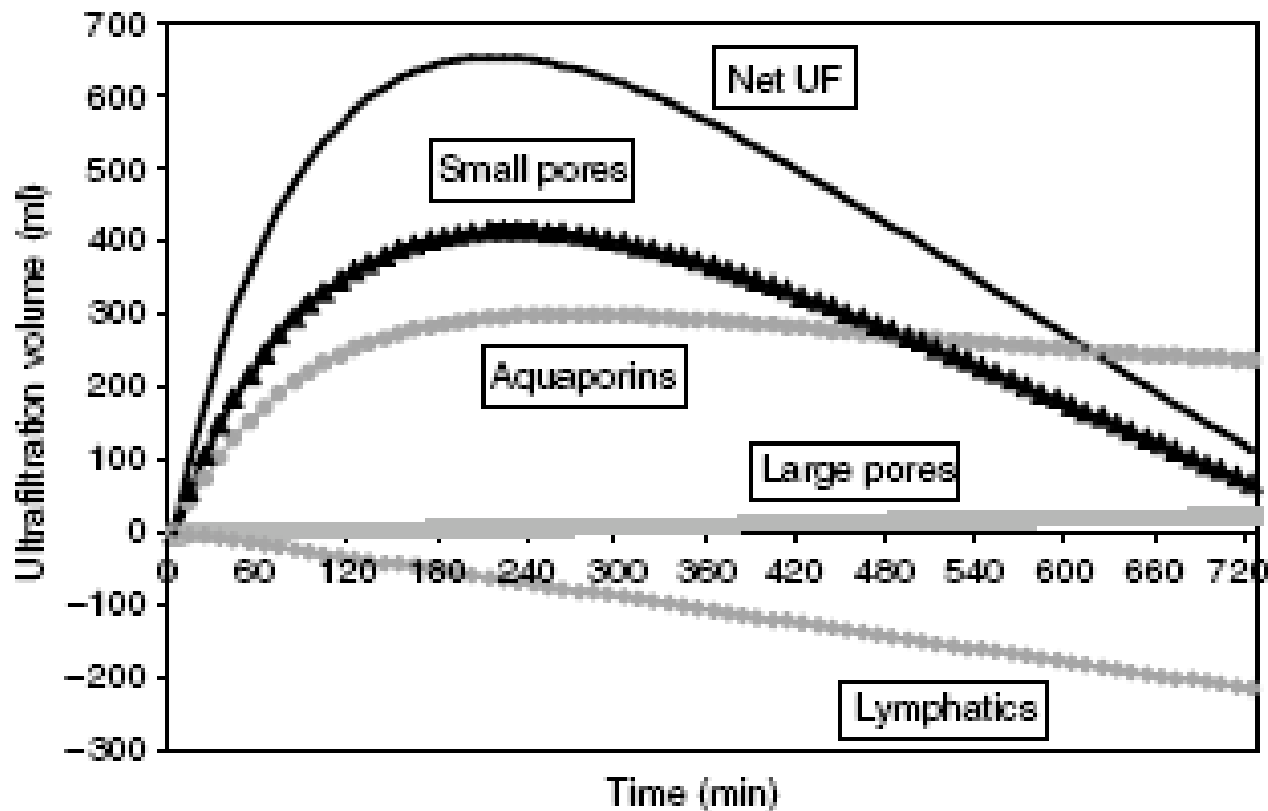
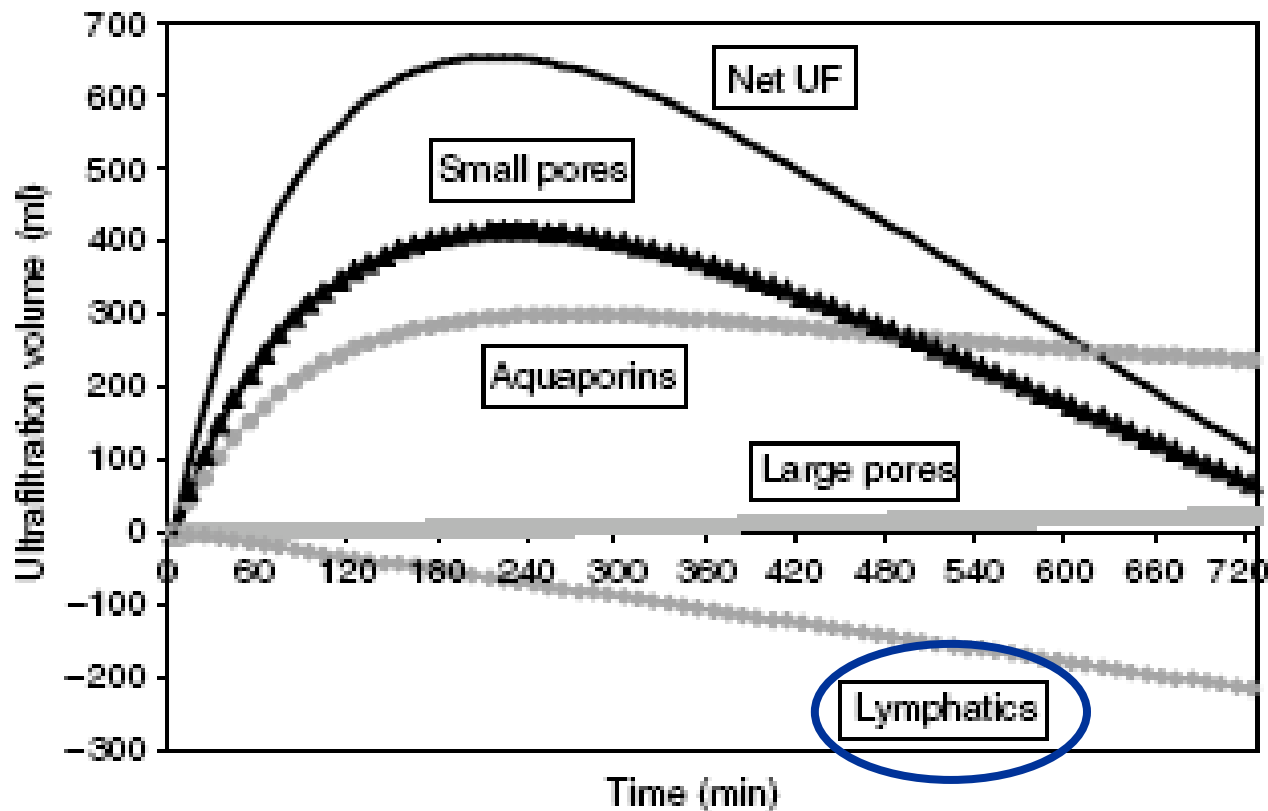


Fig. 1. Schematic model representing CHIP integral membrane protein within the membrane lipid bilayer. Notable features include 1) homotetrameric complex with 1 subunit bearing a polylactosaminoglycan, 2) minimal polypeptide mass extending above or below the lipid bilayer, and 3) possible individual water pore within each subunit.

ULTRAFILTRATION

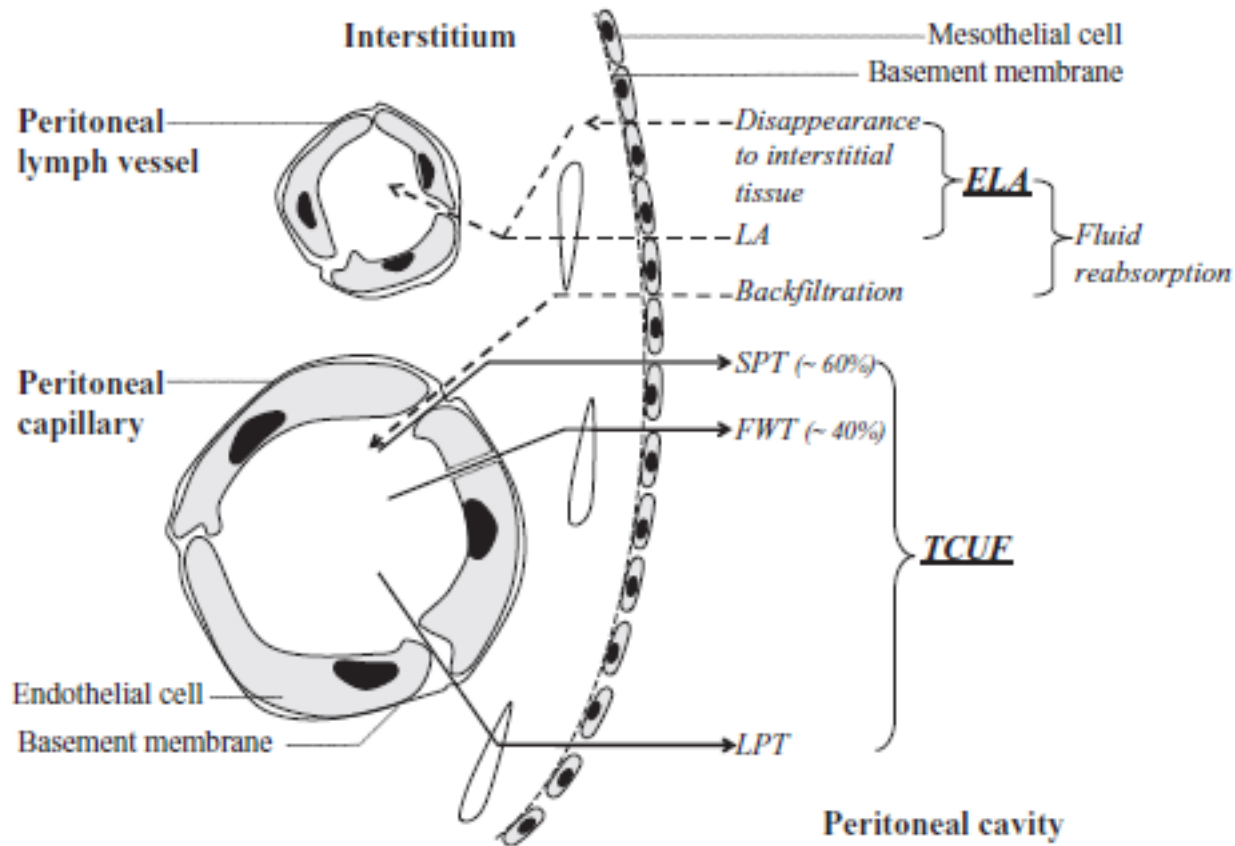


ULTRAFILTRATION

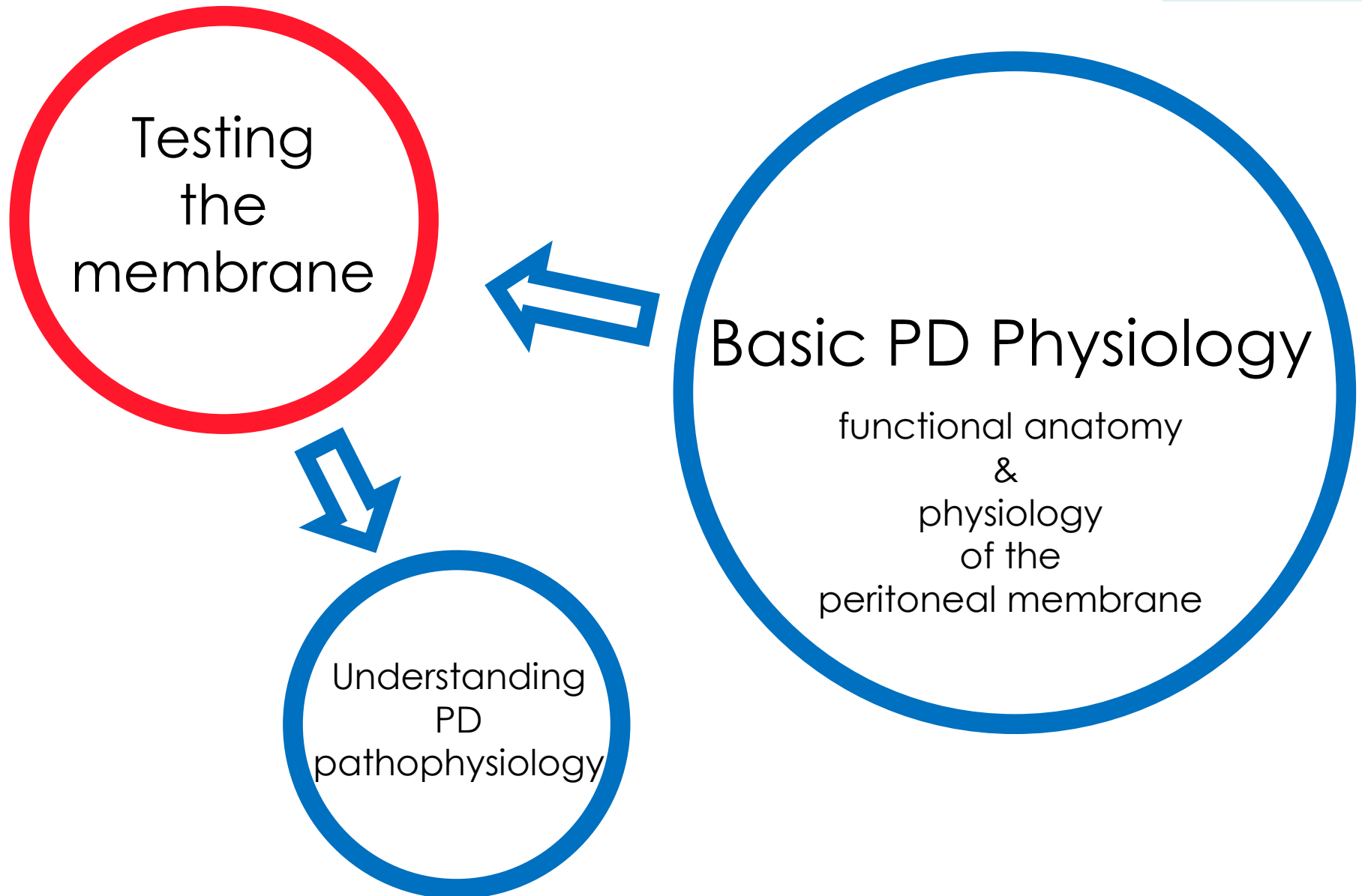




Ultrafiltration



$$NUF = \Delta IPV = TCUF - ELA$$





Peritoneal membrane evaluation

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Peritoneal membrane evaluation



CARI Guidelines 2005



**Canadian Society of Nephrology/
Soci t  Canadienne De N phrologie**

CSN/SCN

Canadian Society of Nephrology, PDI 31:218-239, 2011



National Kidney Foundation™

NKF K/DOQI Am J Kidney Dis 48 (Suppl 1): S138-S142, 2006



UK Renal Association, Guidelines 2010 (review due 2013)



Peritoneal membrane evaluation

ISPD GUIDELINES/RECOMMENDATIONS

GUIDELINE ON TARGETS FOR SOLUTE AND FLUID REMOVAL IN ADULT PATIENTS ON CHRONIC PERITONEAL DIALYSIS



SOLUTE REMOVAL

3. For small solute removal, the total (renal + peritoneal) Kt/V urea should not be less than 1.7 at any time (*Evidence level A*). That means, in anuric patients, peritoneal Kt/V urea has to be above 1.7. In the presence of residual renal function, the contributions of renal and peritoneal clearances may be added for practical purposes, although, as mentioned previously, renal and peritoneal clearances may not be truly additive (*Opinion*). Solute removal above this level should not be equated with “adequate dialysis.” Knowledge of the transport characteristics of the patient’s peritoneal membrane by peritoneal equilibration test or other tests may help to optimize the prescription to meet this target.



ISPD GUIDELINES/RECOMMENDATIONS



EVALUATION AND MANAGEMENT OF ULTRAFILTRATION PROBLEMS IN PERITONEAL DIALYSIS

ULTRAFILTRATION

Recommendations:

- Adherence to sound physiologic principles in the design and implementation of PD prescriptions is essential to prevent the emergence of fluid overload. The most frequently ignored principles in PD that lead to UF difficulties, are the need to avoid long dwells in high transporters, and balancing glucose concentration and dwell time. Prescription setting must take these into account.



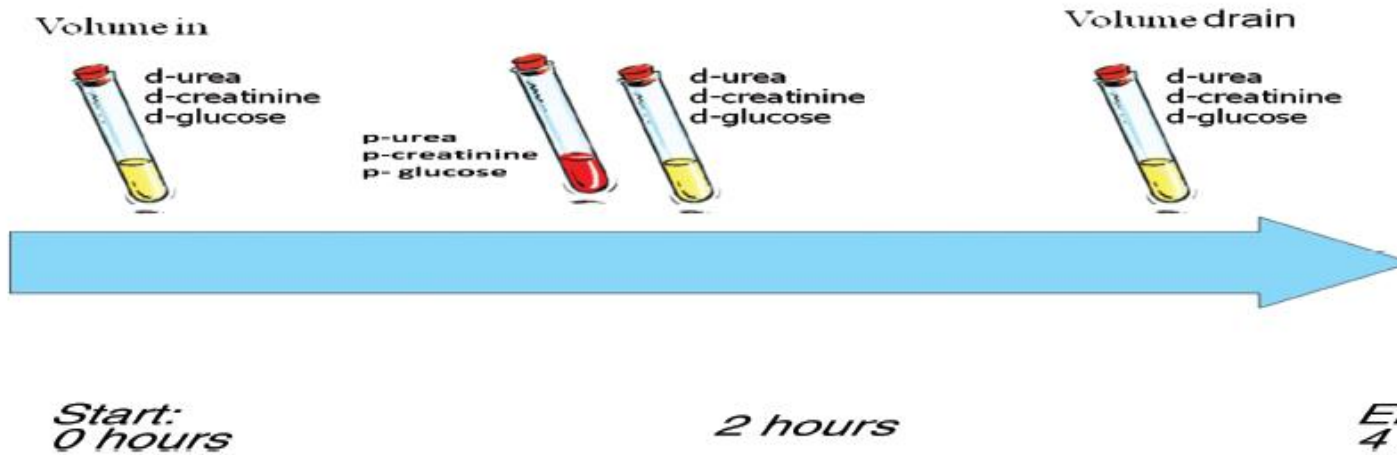
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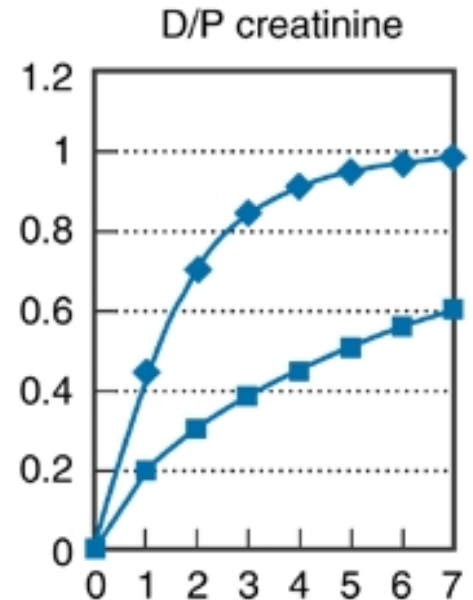


The original 2.27% PET test



SOLUTE TRANSPORT

(D/P creatinine reflects effective vascular surface area, rather than the intrinsic permeability of the membrane!)

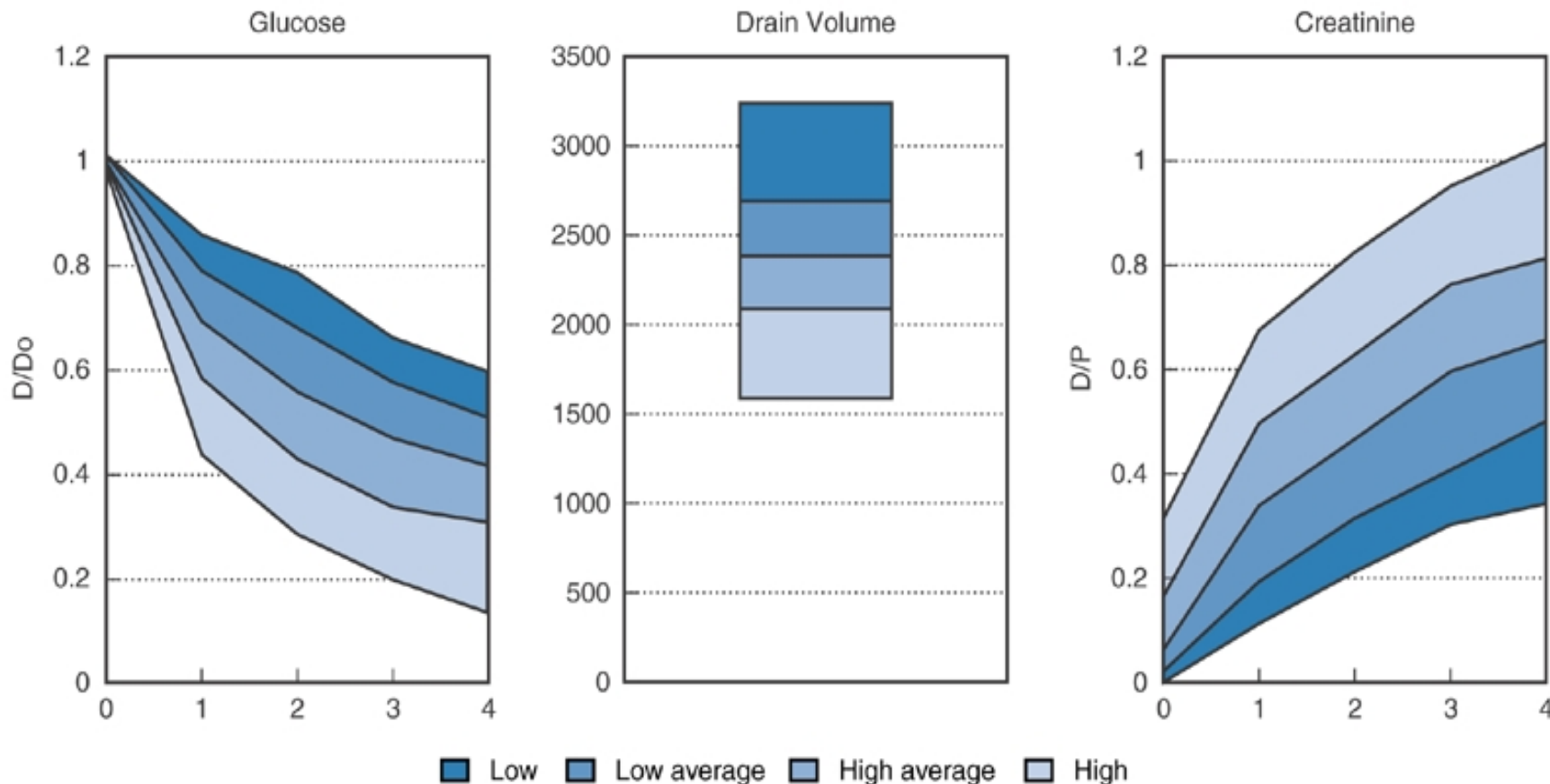




The original 2.27% PET test

SOLUTE TRANSPORT

PERITONEAL EQUILIBRATION TEST



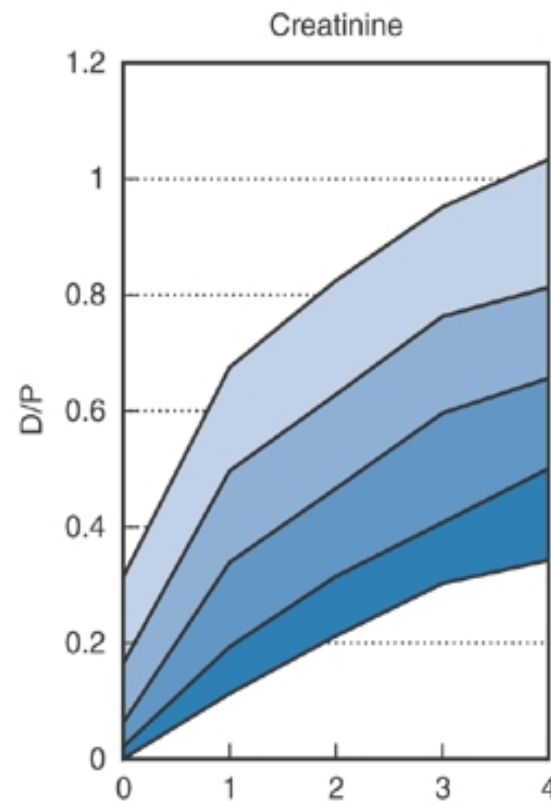
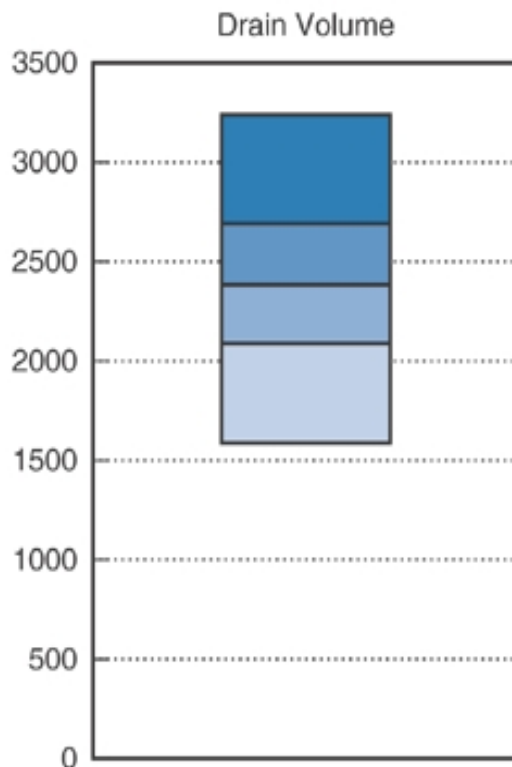
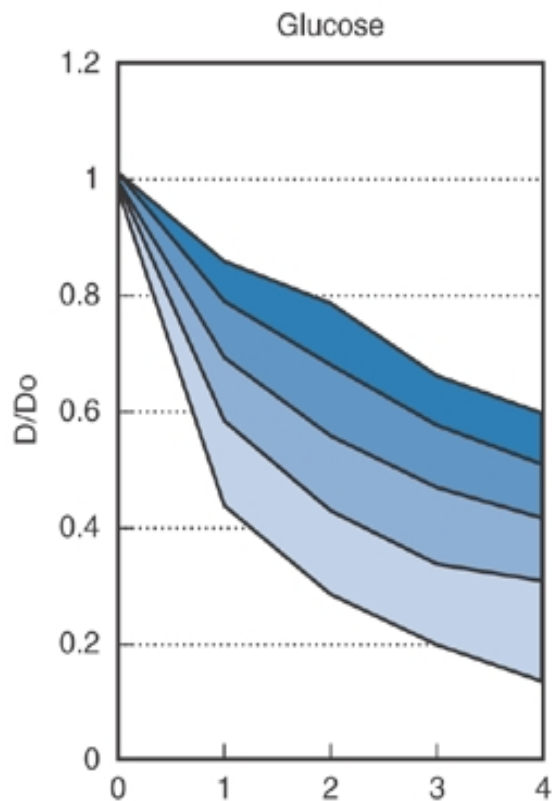


The original 2.27% PET test

ULTRAFILTRATION

SOLUTE TRANSPORT

PERITONEAL EQUILIBRATION TEST



■ Low ■ Low average ■ High average ■ High



The original 2.27% PET test

ULTRAFILTRATION

SOLUTE TRANSPORT

PERITONEAL EQUILIBRATION TEST

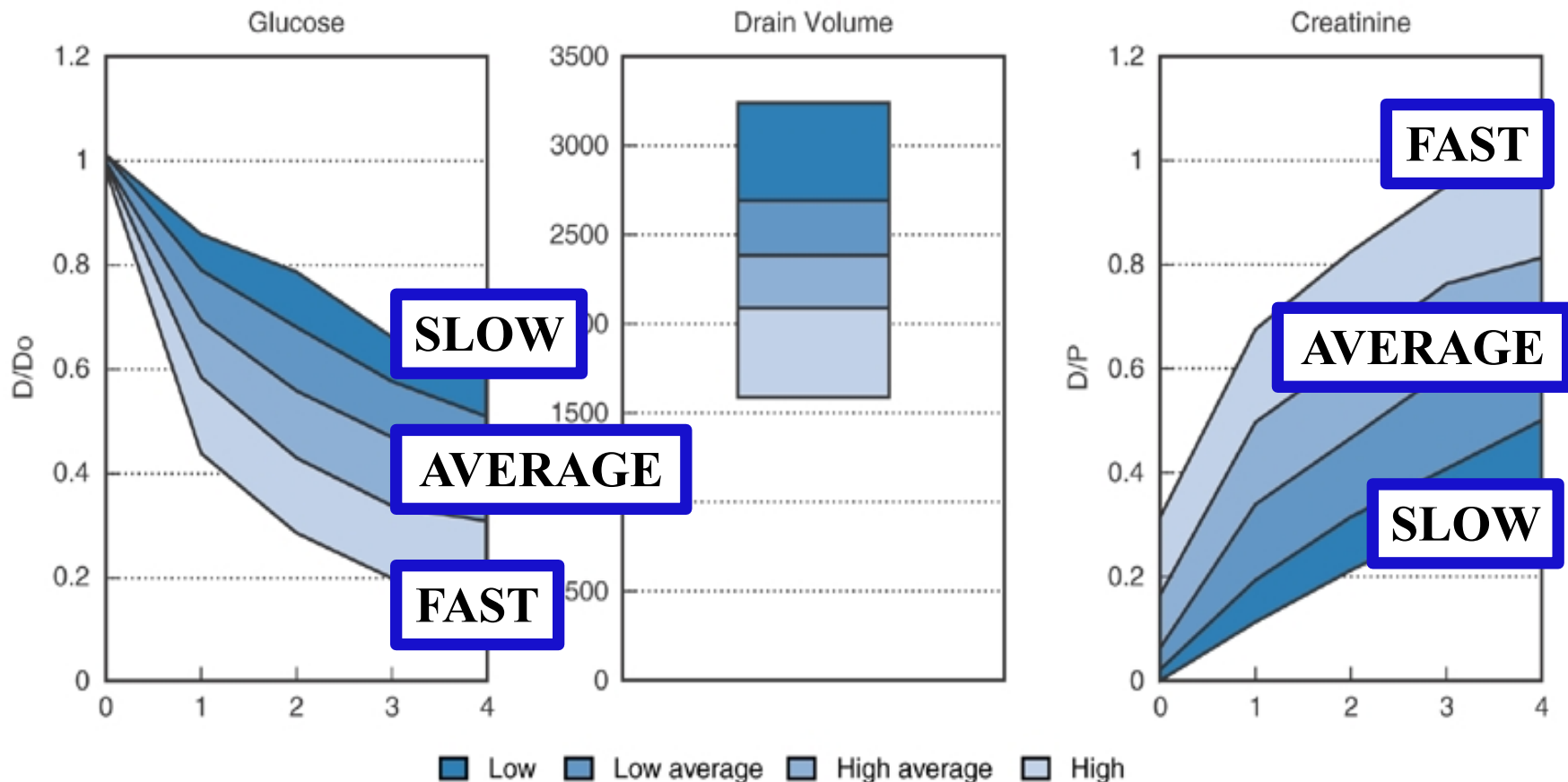
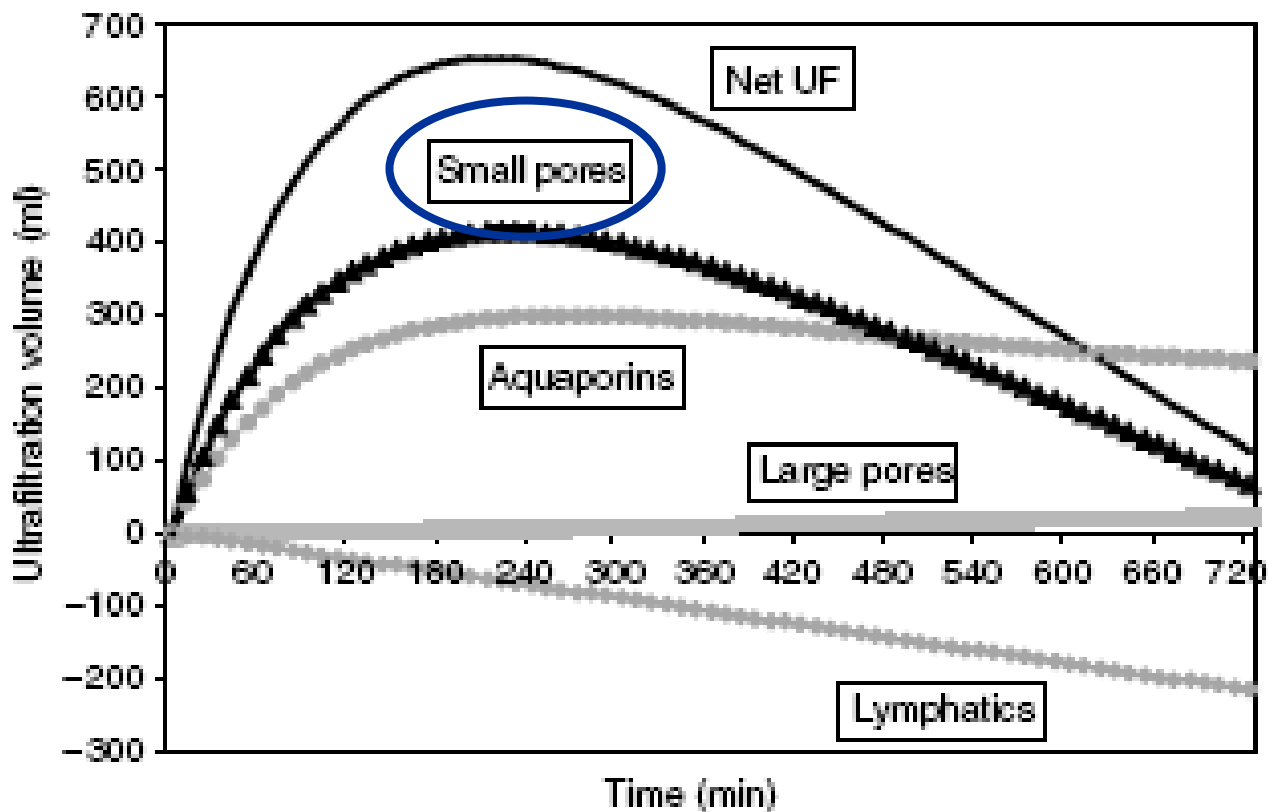


Table 1. Peritoneal membrane transport types and their consequences for clinical management

Transport type	Properties	Recommendations
FAST Fast transporter	Fast, hyperbolic, equilibration of creatinine, typically with a $D/P_{\text{creat}} > 0.80$ after 4 h Fast dissipation of glucose from the peritoneal cavity, with negative ultrafiltration in dwells with 1.36% glucose longer than 180 min Limited sodium sieving, with 3.86% PET and small (< 5 mmol/l) ΔD_{sodium} (difference between the D_{sodium} at start and after 1 h)	Short dwells, preferably shorter than 180 min Icodextrin to be considered for longest dwell, unless sufficient residual diuresis Check inflammatory status (peritoneal protein loss). When negative, check transport status using larger fill volumes
AVERAGE Average transporter	Moderately fast equilibration of creatinine, with a steeper slope in the beginning than at the end of the dwell Moderately fast disappearance of osmotic agent. Negative ultrafiltration only in too long dwells (> 240 min)	Too short (< 120 min) and too long dwells (> 300 min) should be avoided, except for one exchange/day (the 'long dwell')
SLOW Slow transporter	Slow, semi-linear equilibration of creatinine, typically with a $D/P_{\text{creat}} < 0.55-0.60$ after 4 h Sustained ultrafiltration even in dwells longer than 240 min Important sodium sieving, with 3.86%-PET and substantial $\Delta D_{\text{sodium}} (> 5$ mmol/l) after 1 h (the peak of ΔD_{sodium} could occur later in the dwell)	Long dwells, preferably longer than 240 min Use larger volumes rather than more dwells Icodextrin probably not necessary for longest dwell Be aware of sodium sieving when using dwells shorter than 180 min

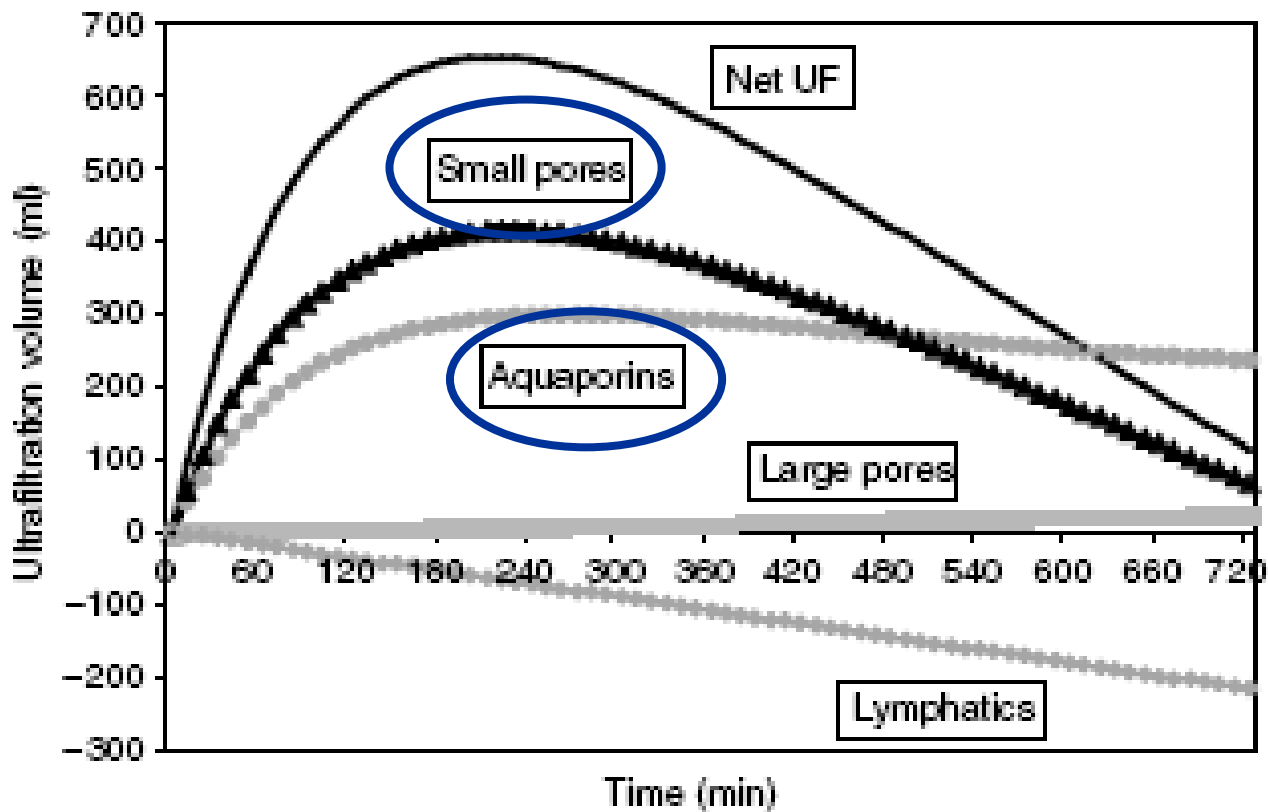
Can be derived from clinical observation without need of formal testing!

ULTRAFILTRATION



The aquaporins?

ULTRAFILTRATION

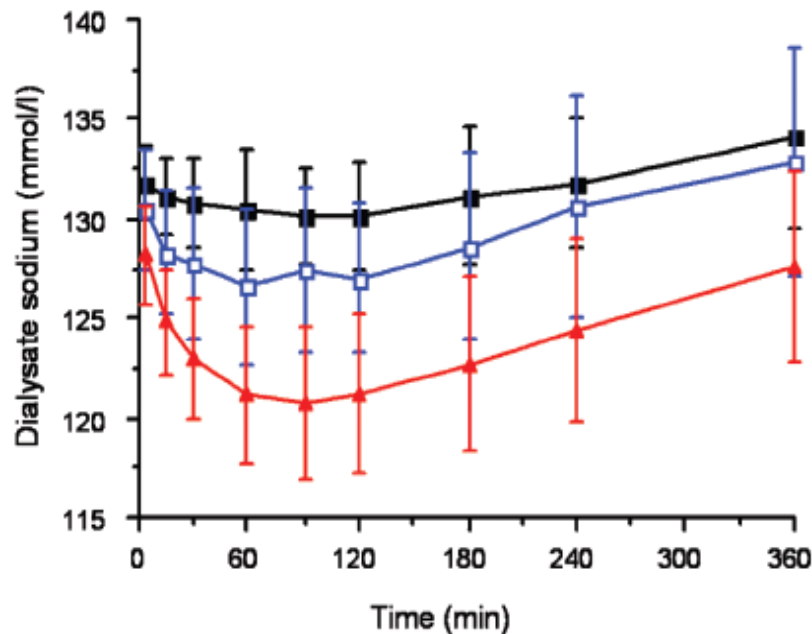


With a hypertonic dialysate solution, dialysate Na⁺ concentration will decrease initially due to water-only transport across aquaporins.

= SODIUM SIEVING

Time profile D/P_{sodium} , D_{sodium} (or D/D_0 or ΔD_{sodium} at 1 hour) CAN BE USED TO ASSESS THE CONTRIBUTION OF AQUAPORIN TRANSPORT TO ULTRAFILTRATION

ISPD definition of UF failure = < 400ml UF after 4 hours of 3.86% glucose

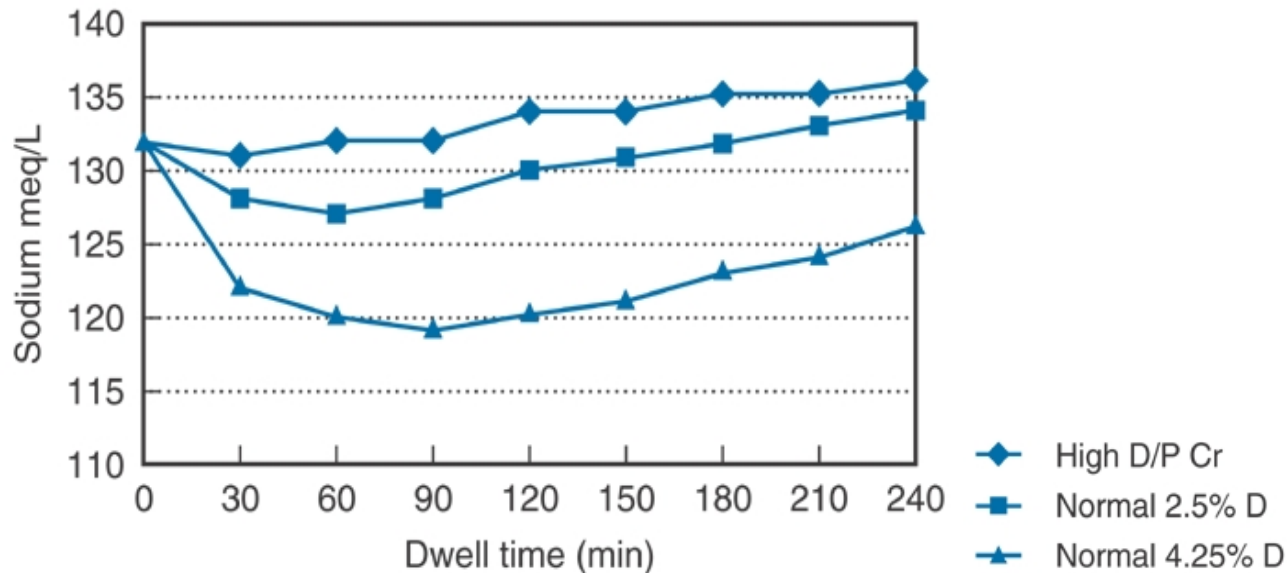


Black: 1.36%; blue: 2.27%; red: 3.86% glucose solution

BUT:
A flat SODIUM SIEVING profile may have different meanings!
(at least theoretically)

aquaporin deficiency

“very very fast” small solute transport (small pores)



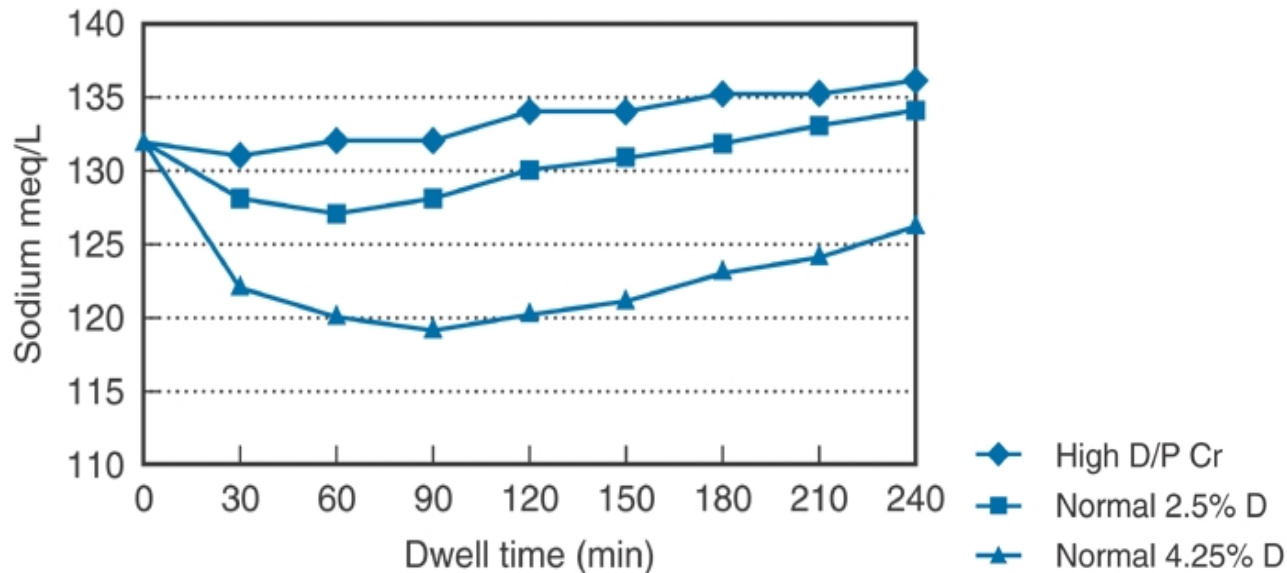
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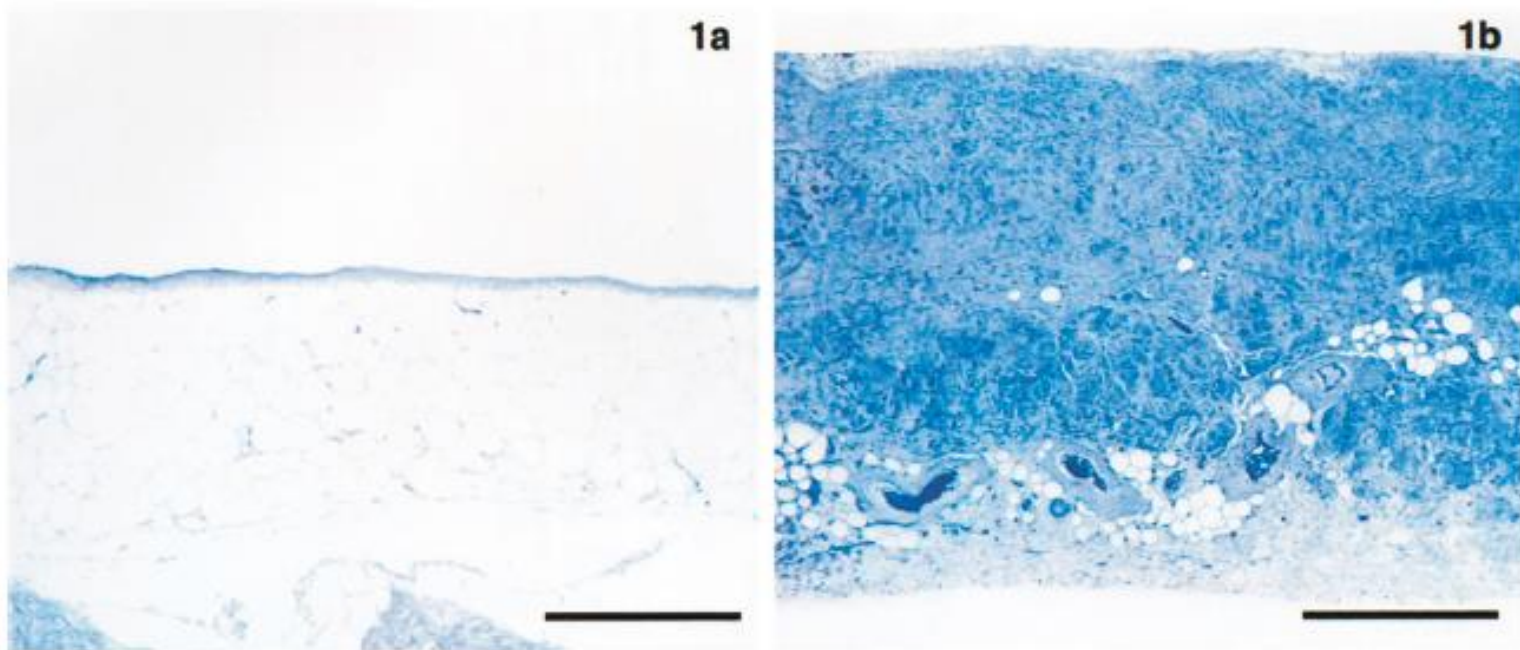
aquaporin deficiency

“very very fast” small solute transport (small pores)

fibrotic peritoneal interstitium (“closed membrane”, uncoupling)



Morphological changes in peritoneal membrane THICKNESS OF SUBMESOTHELIAL COMPACT ZONE



Normal

After 9 years of PD



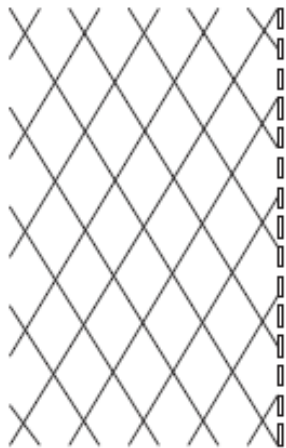
Pore models: interstitium?

the serial three-pore membrane/fiber matrix model

A Three pore membrane with a normal
("loose") serial fiber matrix

$$\mathcal{E} = 0.995$$
$$r_f = 6 \text{ (\AA)}$$

$L_p S \sigma_g = 3.66$	$\mu\text{L}/\text{min}/\text{mmHg}$
$PS_g = 9.30$	mL/min
$\sigma_g = 0.047$	
$L_p S = 0.078$	$\text{mL}/\text{min}/\text{mmHg}$

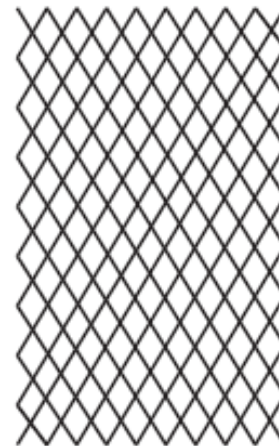


$S = 1$

B Three pore membrane with a fibrotic
("dense") serial fiber matrix

$$\mathcal{E} = 0.96$$
$$r_f = 7.5 \text{ (\AA)}$$

$L_p S \sigma_g = 3.02$	$\mu\text{L}/\text{min}/\text{mmHg}$
$PS_g = 13.46$	mL/min
$\sigma_g = 0.039$	
$L_p S = 0.078$	$\text{mL}/\text{min}/\text{mmHg}$



$S = 1.8$

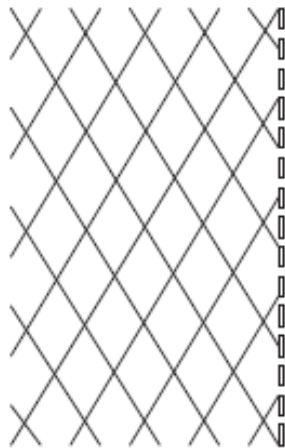
the serial three-pore membrane/fiber matrix model

A Three pore membrane with a normal ("loose") serial fiber matrix

$$\mathcal{E} = 0.995$$

$$r_f = 6 \text{ (Å)}$$

$L_p S \sigma_g$	= 3.66	$\mu\text{L}/\text{min}/\text{mmHg}$
PS_g	= 9.30	mL/min
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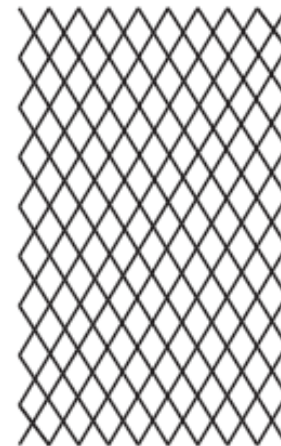
$S = 1$

B Three pore membrane with a fibrotic ("dense") serial fiber matrix

$$\mathcal{E} = 0.96$$

$$r_f = 7.5 \text{ (Å)}$$

$L_p S \sigma_g$	= 3.02	$\mu\text{L}/\text{min}/\text{mmHg}$
PS_g	= 13.46	mL/min
σ_g	= 0.039	
$L_p S$	= 0.078	$\text{mL}/\text{min}/\text{mmHg}$



$S = 1.8$

The Osmotic Conductance to Glucose

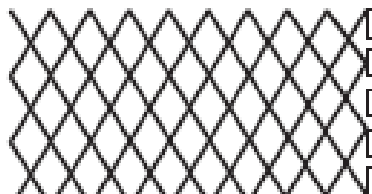
= the ability of glucose to exert an osmotic pressure sufficient to cause transperitoneal ultrafiltration

$$= L_p \cdot S \cdot \sigma \text{ (}\mu\text{L/min/mmHg)}$$

B Three pore membrane with a fibrotic ("dense") serial fiber matrix

$$\begin{aligned} \mathcal{E} &= 0.96 \\ r_f &= 7.5 \text{ (}\AA\text{)} \end{aligned}$$

$L_p S \sigma_g$	= 3.02	$\mu\text{L/min/mmHg}$
PS_g	= 13.46	mL/min
σ_g	= 0.039	
$L_p S$	= 0.078	mL/min/mmHg





OCG: the Dummy's view

$L_p \cdot S \cdot \sigma$ ($\mu\text{l}/\text{min}/\text{mmHg}$)

Reflection coefficient of glucose

= lower in case of aquaporin dysfunction

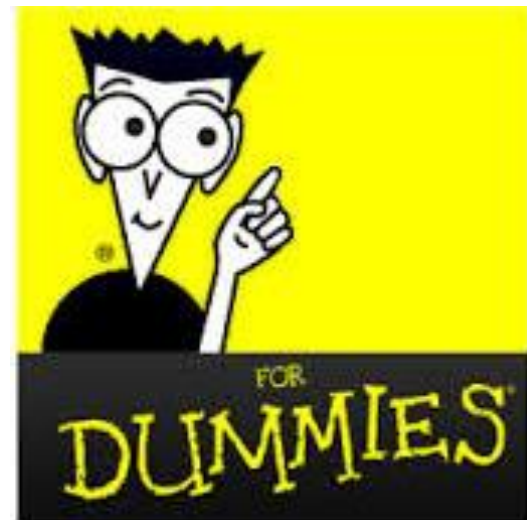
= lower in case of increased small solute transport

Surface area

= higher in case of increased small solute transport

Hydraulic conductivity

= lower in case of fibrosis



**A flat SODIUM SIEVING profile may have different meanings!
(at least theoretically)**

$L_p \cdot S \cdot \sigma$ ($\mu\text{l}/\text{min}/\text{mmHg}$)

aquaporin deficiency

“very very fast” small solute transport (small pores)

fibrotic peritoneal interstitium (“closed membrane”, uncoupling)

	OCG	Free water transport	Small pore water transport
Reference	normal	normal	normal
Increased small solute transport	normal	normal	low
Aquaporin dysfunction	low	low	normal
Fibrotic interstitium	low	low	low

A flat SODIUM SIEVING profile may have different meanings! (at least theoretically)

aquaporin deficiency

“very very fast” small solute transport (small pores)

fibrotic peritoneal interstitium (“closed membrane”, uncoupling)

Mini-PET added value:

**quantitative assessment of free water transport
and small pore water transport**

Double mini-PET added value:

**quantitative assessment of free water transport
and small pore water transport**

+

assessment of osmotic conductance to glucose



(double) mini-PET

Mini-PET

1 hour of 3.86%

**assessment of $D/P_{\text{creatinin}}$, D/P_{glucose} , D/D_0 or ΔD_{sodium}
calculation of free & small pore water transport**

Free water transport (FWT):

$$\text{FWT (ml)} = \text{UFT (ml)} - \text{UFSP (ml)}$$

Ultrafiltration over the small pores (UFSP) is assessed using the Na clearance:

$$\text{UFSP (ml)} = [\text{NaR (mmol)} 1000] / \text{Na}_p \text{ (mmol/l)}$$

NaR (mmol) is the Na removed during the second part of the test with the 3.86% solution. NaR is calculated as follows:

$$\text{NaR (mmol)} = [\text{Drained dialysate volume (l)} \cdot \text{Na concentration (mmol/l) in the drained dialysate}] - [\text{Volume of dialysate before infusion (l)} \cdot \text{Na concentration (mmol/l) in dialysate before infusion}]$$

$\text{Na}_p = \text{plasma sodium.}$



(double) mini-PET

Mini-PET

1 hour of 3.86%

assessment of $D/P_{\text{creatinin}}$, D/P_{glucose} , D/D_0 or ΔD_{sodium}

calculation of free & small pore water transport

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Na_p = plasma sodium.

Double mini-PET

1 hour of 1.36%, followed by 1 hour of 3.86%

From the 3.86% hour:

assessment of $D/P_{\text{creatinin}}$, D/P_{glucose} , D/D_0 or ΔD_{sodium}
calculation of free & small pore water transport

From the two consecutive dwells of 1 hour:

Osmotic Glucose Conductance (OCG)
(ml/min/mmHg)

$$OCG = ((V_{3.86} - V_{1.36}) / (19.3(G_{3.86} - G_{1.36})60))1.7$$



(double) mini-PET \longrightarrow uni-PET



Double mini-PET

**$D/P_{\text{creatinin}}$, D/P_{glucose}
cannot be extrapolated to the classical or modified PET result**

**Ultrafiltration volume after 1 hour 3.86%
cannot be referenced to the ISPD definition of UF failure**

Rodriguez et al. Blood Purif 25: 497-504, 2007
Waniewski et al. ASAJO J 42: 518-523, 1998
Imholtz et al. Kidney Int 46: 333-340, 1994



(double) mini-PET → uni-PET

Uni-PET

**1 hour of 1.36%,
followed by 4 hours of 3.86%,
but with temporary drainage
after 1 hour**

UNI-PET		naam:	datum:			
Concentratie dialysaat	Tijd	Procedure UNI-PET	volume In	volume uit	naam dialysaat staal uit	bloed staal
2,27% 2l	s' avonds	. wegen zak+ emmer+klemmen . 8-10u terplaatse		in dagboek pat		
Na 8-10 u verblijftijd 2,27 %						
1,36%2L	TO	. wegen 1,36% zak +emmer+klemmen . rechtopstaande uitloop over 20min . inloop over 10 min, liggende houding rotatie om 400ml . dialysaatstaal direct na inloop 1,36% (100ml uit, staalname en reinfusion) . wegen uitloop nacht 60 min verblijftijd start na inloop			Dt-60	
Na 60 min verblijftijd 1,36%						
3,86%2l	TO	. wegen 3,86% zak+emmer+klemmen . uitloop 1,36% over 20 min . Inloop 3,86% over 10min . dialysaatstaal uitloop 1,36% . dialysaatstaal direct na inloop 3,86% (100 ml uit, staalname en reinfusion) . Wegen uitloop 1,36% 60 min verblijftijd start na inloop			Dref DT0	
Na 60 min verblijftijd 3,86 %						
	T60	. volledige uitloop over 20min in enkelvoudige uitloopzak . wegen uitloop . uitloop terug laten inlopen, 10ml in zak laten voor staalname dialysaat . bloedstaal 120 min verblijftijd start na inloop			DT60	PT60
Na 120min verblijftijd 3,86 %						
	T120	. dialysaatstaal (100ml uit, staalname en reinfusion)			DT120	
. Na 240 min verblijftijd 3,86%						
dialysaat zie voorschrift	T240	wegen dialysaatzak+emmer+klemmen . uitloop over 20 min . staal uitloop . inloop (zie voorschrift) . wegen uitloop		in dagboek pat		DT240

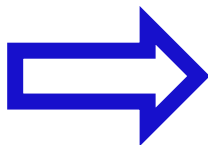
**A flat SODIUM SIEVING profile may have different meanings!
(at least theoretically)**

$$L_p \cdot S \cdot \sigma \text{ (}\mu\text{l/min/mmHg)}$$

aquaporin deficiency

“very very fast” small solute transport (small pores)

fibrotic peritoneal interstitium (“closed membrane”, uncoupling)



	OCG	Free water transport	Small pore water transport
Reference	normal	normal	normal
Increased small solute transport	normal	normal	low
Aquaporin dysfunction	low	low	normal
Fibrotic interstitium	low	low	low



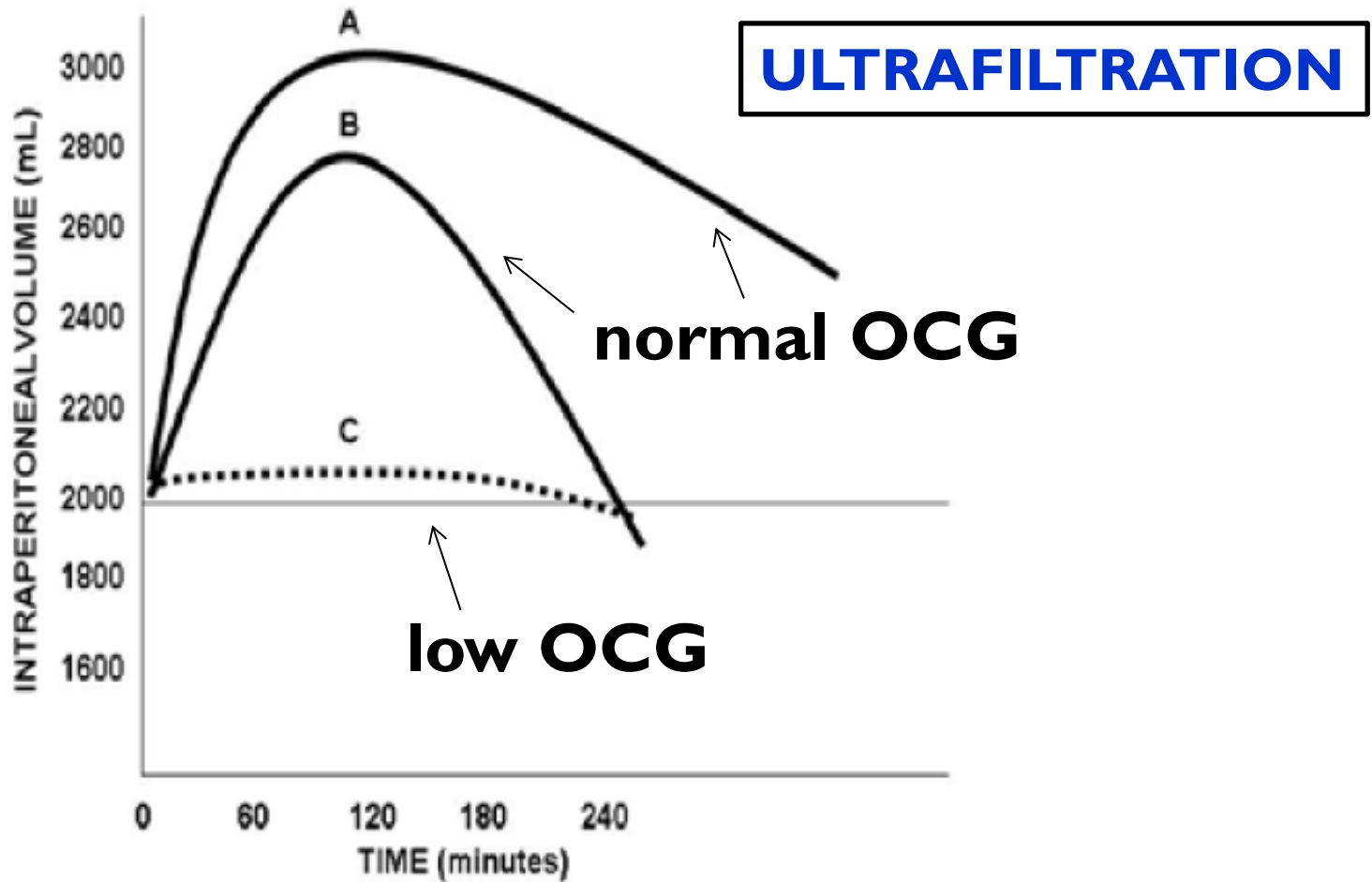
OCG: what does it mean?

Parameters	Transport groups			ANOVA, P
	Slow-average, $n = 6$	Fast-average, $n = 13$	Fast, $n = 5$	
L_pS (mL/min/mmHg)	0.023 ± 0.023	0.030 ± 0.024	0.043 ± 0.046	0.53
σ_G	0.139 ± 0.060	0.117 ± 0.078	0.101 ± 0.055	0.60
OCG (mL/min/mmHg)	0.0022 ± 0.0007	0.0023 ± 0.0008	0.0031 ± 0.0015	0.49

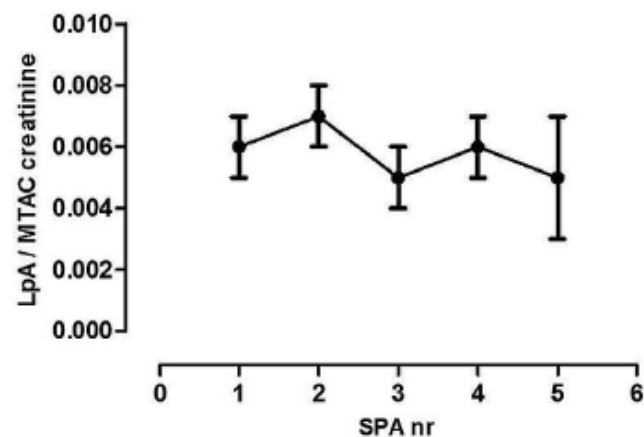
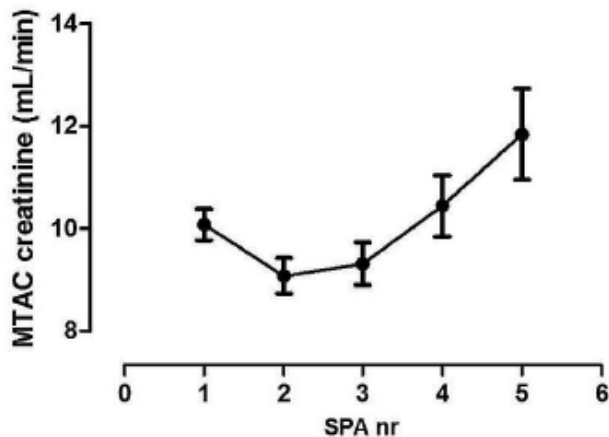
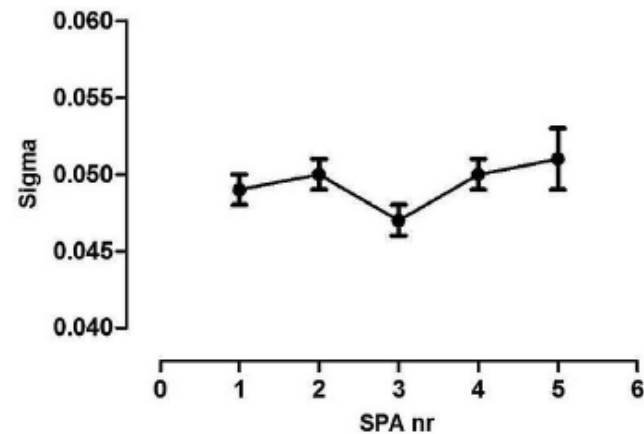
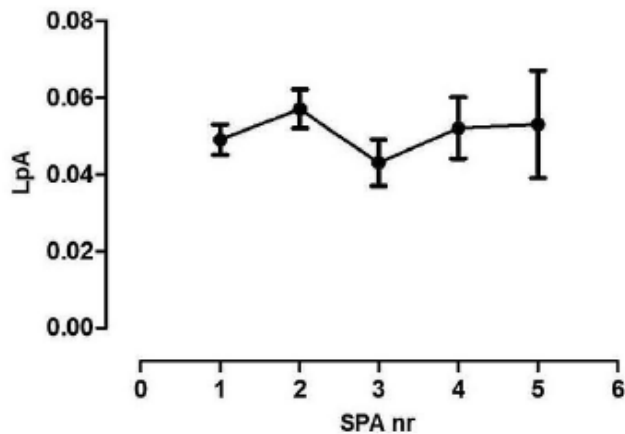
increasing/fast small solute transport
 \neq
low osmotic conductance to glucose

R	nUF60	UFSP60	FWT60	FWF60	nUF240	$A_0/\Delta x$	D/PCr240	D/D0G240	D/PNa60	DipNa60
OCG	0.91 P < 0.001	0.84 P < 0.001	0.11 P = 0.61	-0.50 P < 0.05	-0.09 P = 0.67	0.24 P = 0.26	0.10 P = 0.65	-0.09 P = 0.67	-0.01 P = 0.97	-0.06 P = 0.77

OCG: what does it mean?



Low osmotic conductance to glucose is particularly seen in late ultrafiltration failure.



**A flat SODIUM SIEVING profile may have different meanings!
(at least theoretically)**

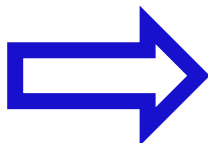
$$L_p \cdot S \cdot \sigma \text{ (}\mu\text{l/min/mmHg)}$$

aquaporin deficiency

“very very fast” small solute transport (small pores)

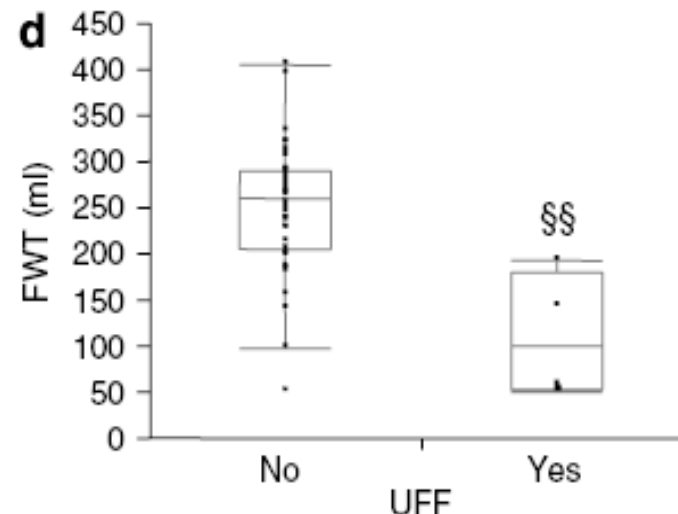
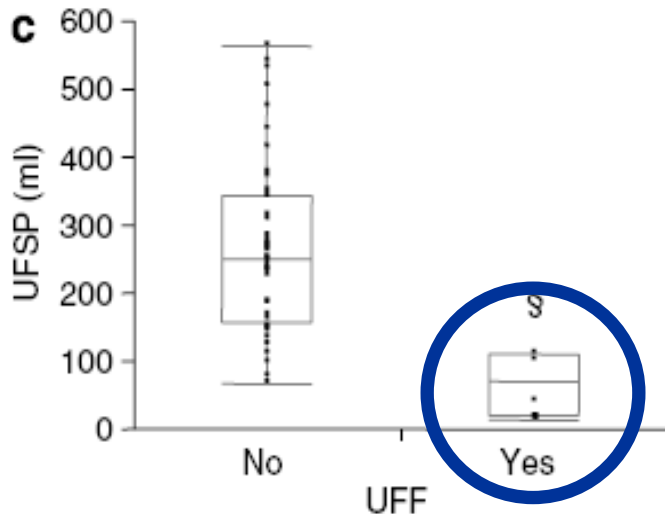
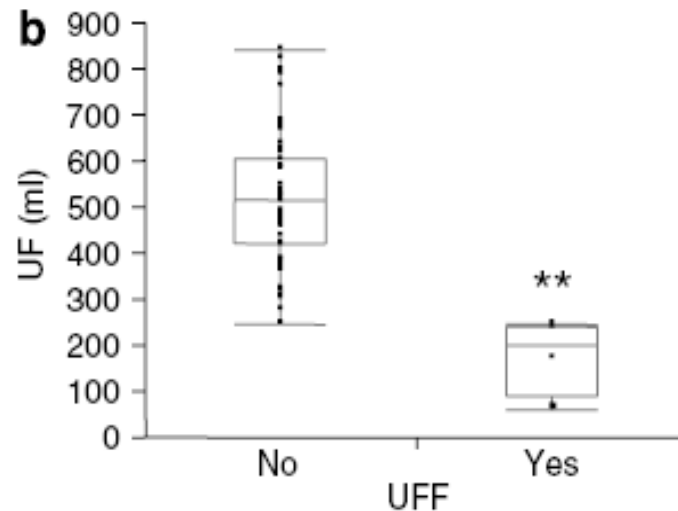
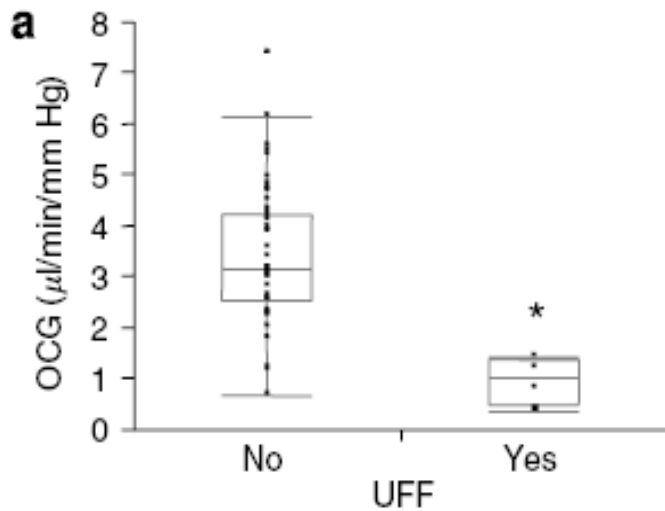
fibrotic peritoneal interstitium (“closed membrane”, uncoupling)

	OCG	Free water transport	Small pore water transport
Reference	normal	normal	normal
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Aquaporin dysfunction	low	low	normal
Fibrotic interstitium	low	low	low



‘isolated aquaporin dysfunction probably non-existent’ (Rippe a.o.)

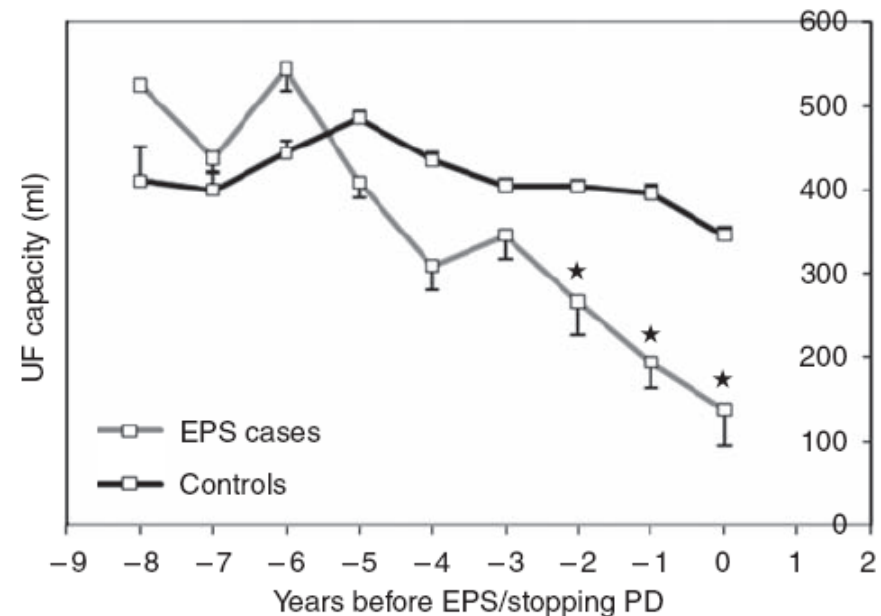
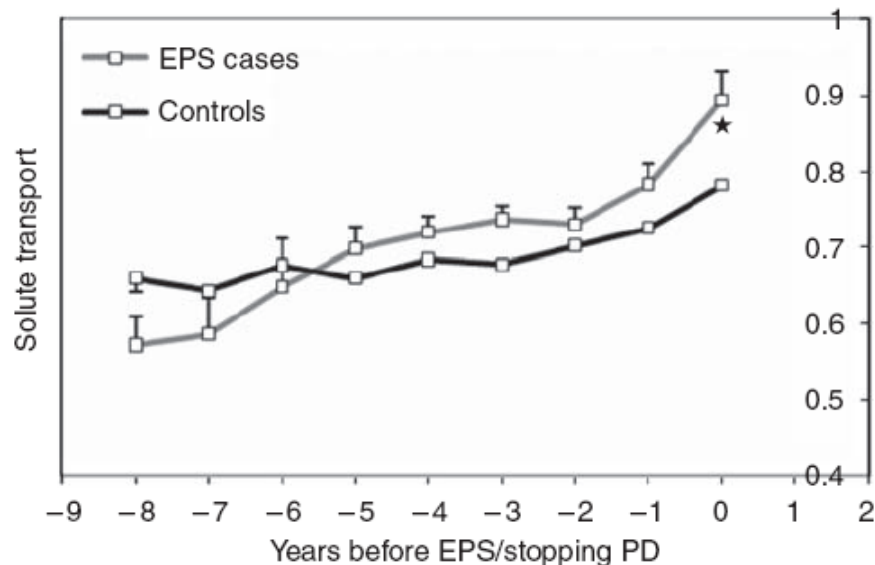
OCG: what does it mean?



Double mini-PET test

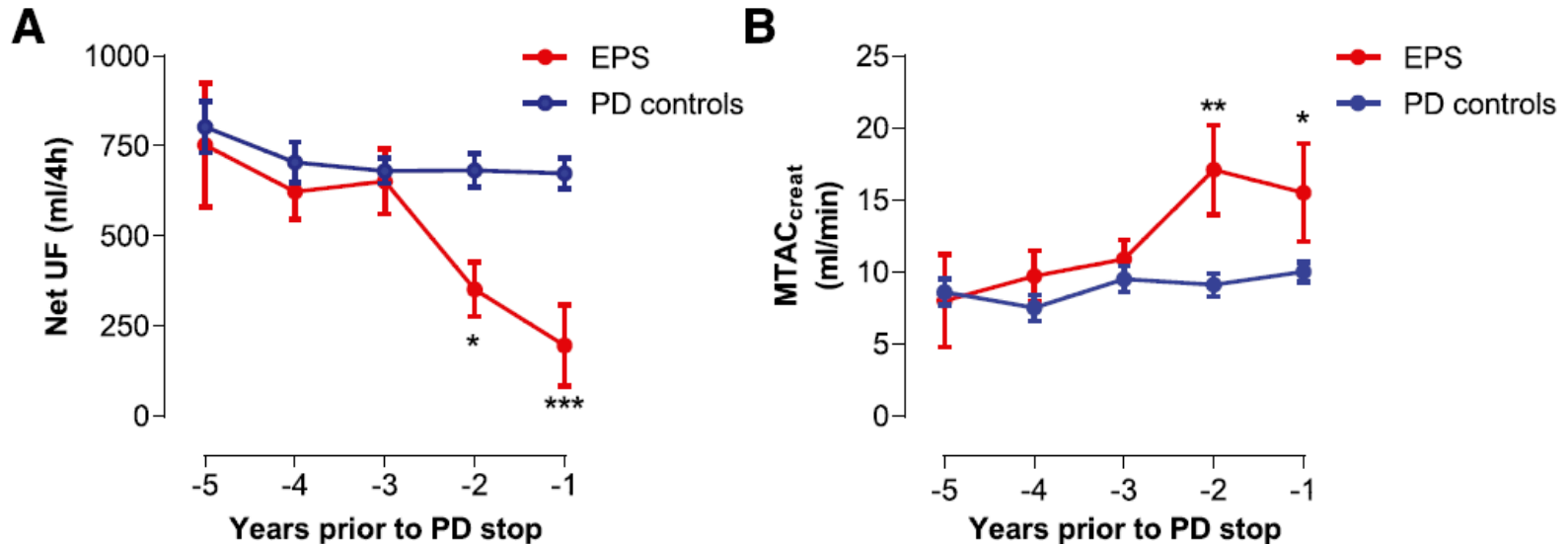
The peritoneal osmotic conductance is low well before the diagnosis of encapsulating peritoneal sclerosis is made

Mark L. Lambie^{1,2}, Biju John^{1,2}, Lily Mushahar^{1,2}, Christopher Huckvale^{1,2} and Simon J. Davies^{1,2}



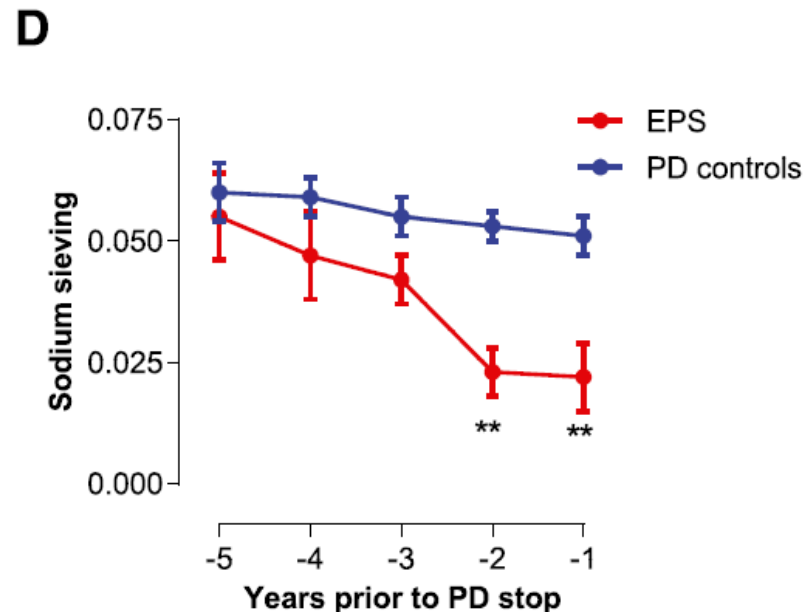
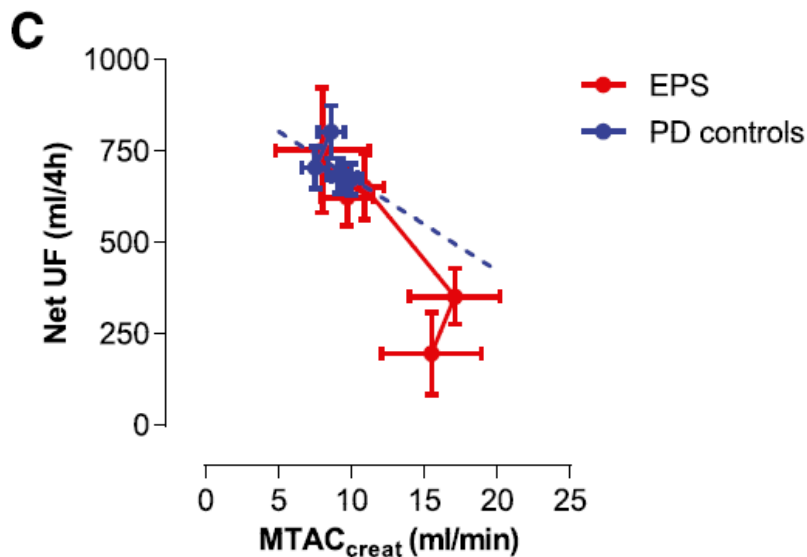
Interstitial Fibrosis Restricts Osmotic Water Transport in Encapsulating Peritoneal Sclerosis

Johann Morelle,^{*} Amadou Sow,^{*} Nicolas Hautem,^{*} Caroline Bouzin,[†] Ralph Crott,[‡] Olivier Devuyst,^{*§} and Eric Goffin^{*}



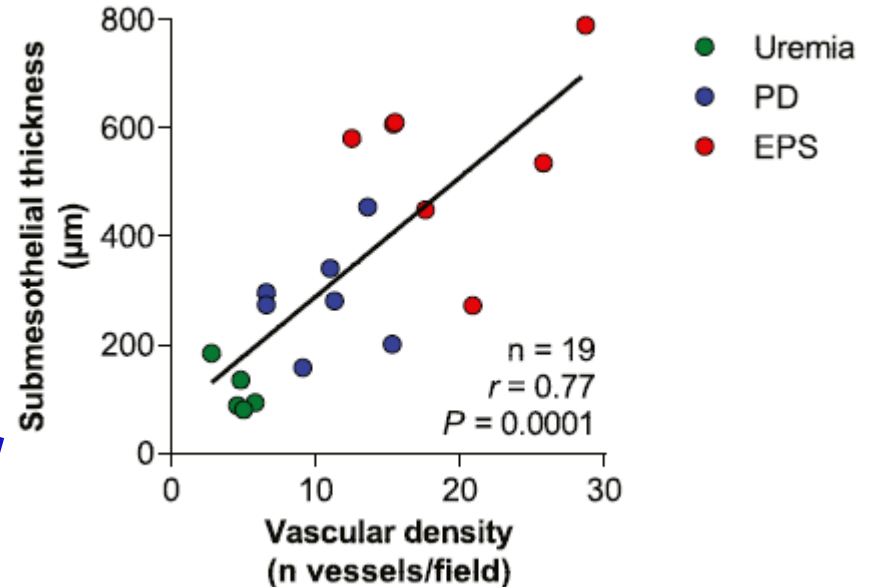
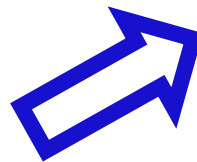
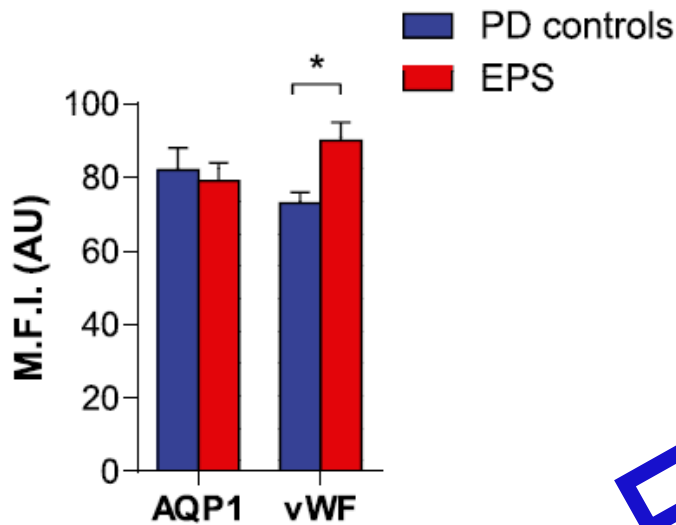
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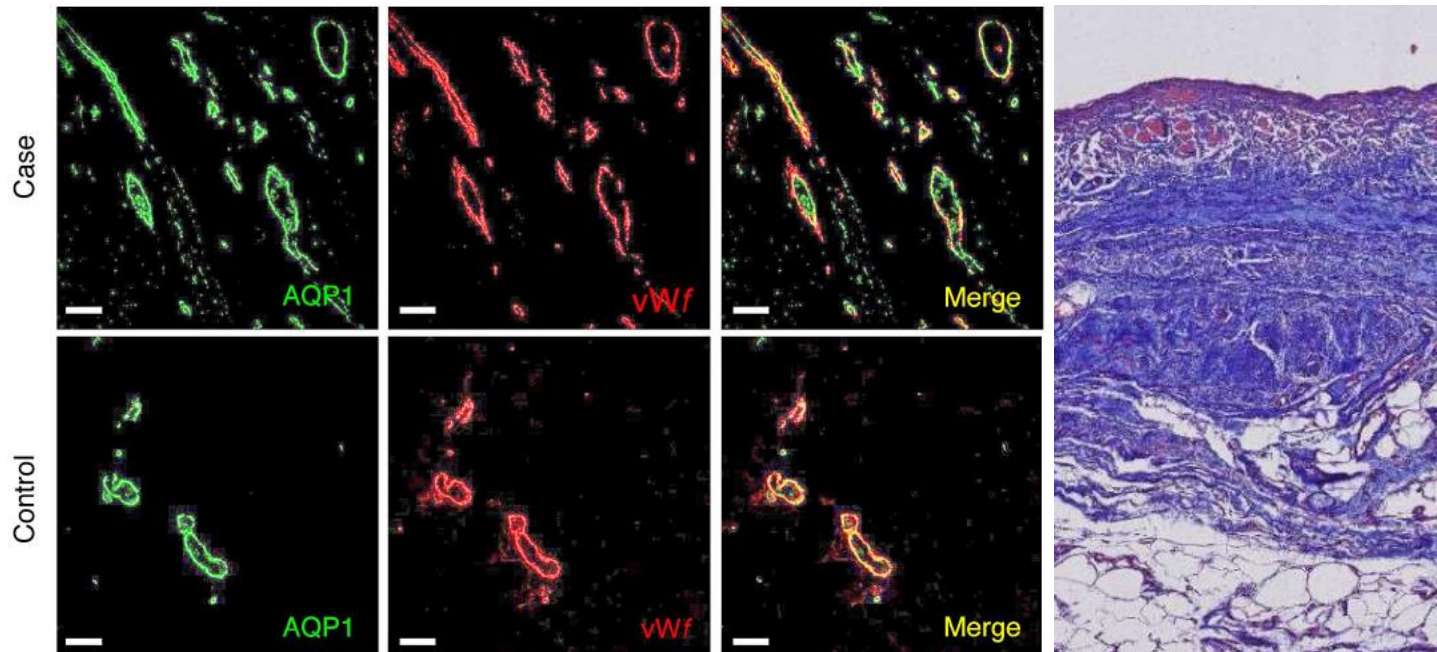
Johann Morelle,^{*} Amadou Sow,^{*} Nicolas Hautem,^{*} Caroline Bouzin,[†] Ralph Crott,[‡] Olivier Devuyst,^{*§} and Eric Goffin^{*}





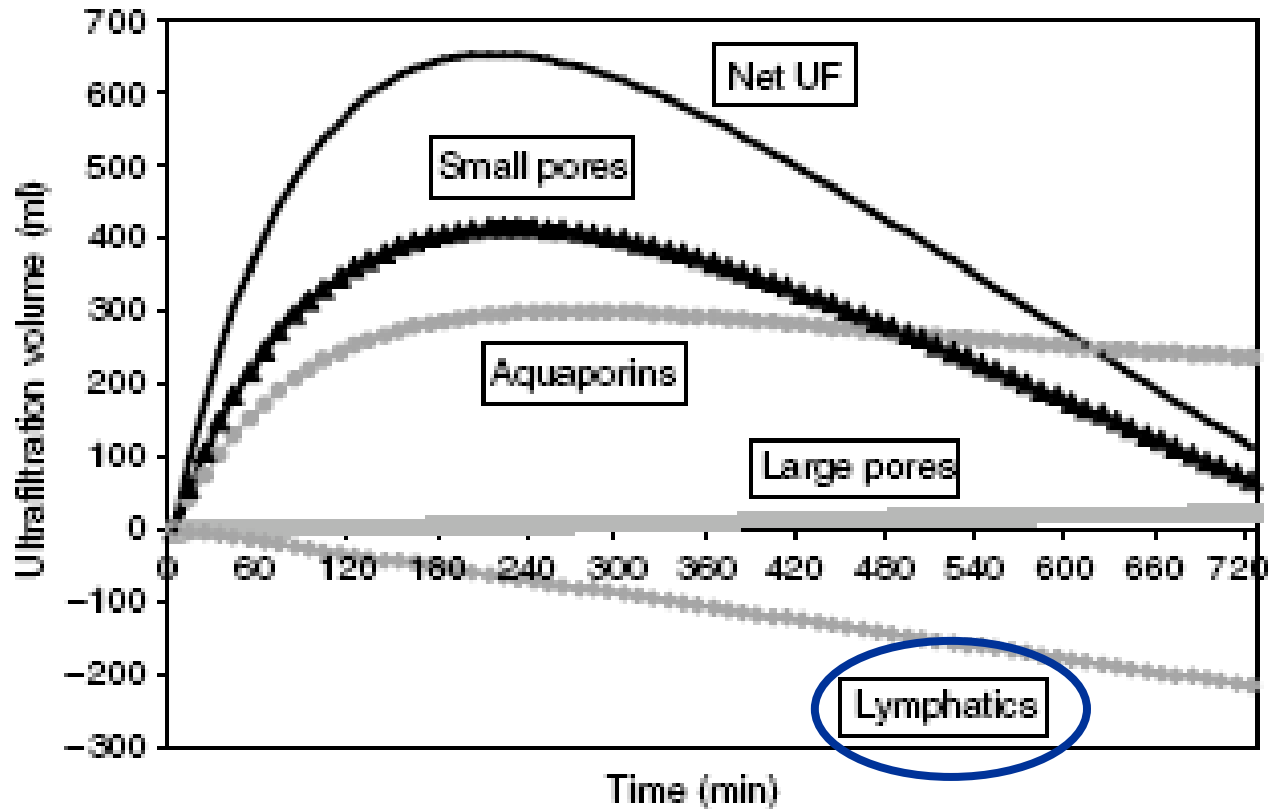
OCG: what does it mean?

Ultrafiltration Failure and Impaired Sodium Sieving During Long-Term Peritoneal Dialysis: More Than Aquaporin Dysfunction?



The THREE pore theory

ULTRAFILTRATION





The standard peritoneal permeability analysis: A tool for the assessment of peritoneal permeability characteristics in CAPD patients

MARJA M. PANNEKEET, ALEXANDER L.T. IMHOLZ, DICK G. STRUIJK, GER C.M. KOOMEN,
MONIQUE J. LANGEDIJK, NATALIE SCHOUTEN, RUDI DE WAART, JOHAN HIRALALL,
and RAYMOND T. KREDIET

1.36% glucose solution + Dextran 70 (1g/L)

allows calculation of

MTAC (i.e. D/P) of creatinin, urea, urate
glucose absorption rate
net ultrafiltration

but also

effective lymphatic absorption rate
clearances of other molecules (β_2 M, IgG,...)

Now you know
how to evaluate the peritoneal membrane!

BUT...

...IS IT NOT USEFUL ANYMORE?

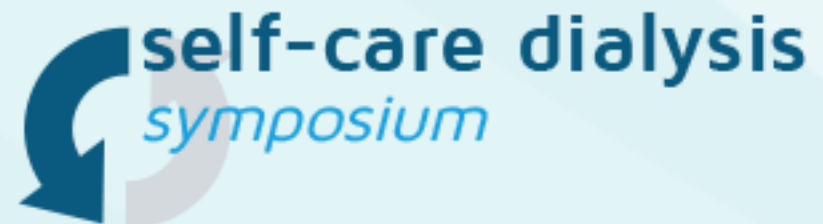
...IS IT STILL MANDATORY?



WHO KNOWS THE ANSWER?



3rd self-care
dialysis symposium
12th & 13th May 2016



How to evaluate the peritoneal membrane?



B. Bammens

Brussels, May 12 2016

UZ
LEUVEN

KU LEUVEN

BELGIUM

3rd self-care
dialysis symposium
12th & 13th May 2016

