Advances in Hemodialysis Therapy & Technology

### Hesham Elsayed

**Emeritus Prof of Nephrology ASU** 

ESNT president elect

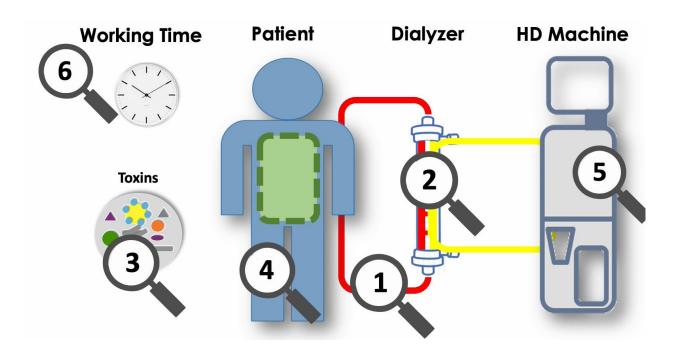
HD chapter chair

HD guideline editor



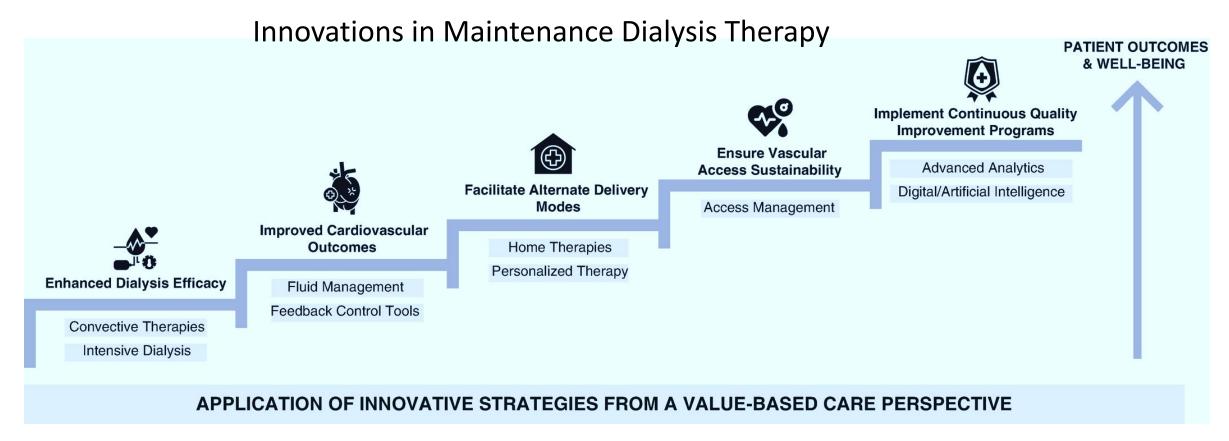
### Agenda

- 1- Patient Centered Approach
- 2- Convection therapies , HDF and HDX
- 3- Extracorporeal therapies in ICU
- 4- Volume and Electrolyte control
- 5- DOACs in AF with ESKD patients
- 6- Cellular activation during HD
- 7- Home Hemodialysis
- 8- Future of intracorporeal HD



# A Gap between HD Adequacy and Patient Centered Needs: A new Definition for improving outcomes





Incremental and collective application of different strategies to target improvement of patient outcomes in terms of reducing morbidity and mortality that is still high for the dialysis population

### The current and future landscape of dialysis

### **Patient priorities**

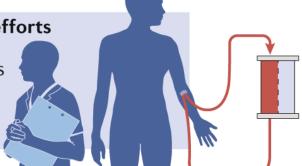
- Physical symptoms (fatigue, insomnia, cramps, pain)
- Mood symptoms (depression, anxiety, frustration, wash-out)
- Rehabilitation priorities (ability to work, ability to travel, impact on family and friends, mobility)

### Top-down efforts (by government agencies, societies, NGOs, etc.)

- Regulatory considerations
- Reimbursement and other financial incentives
- Guidance on product development and clinical end points
- Support for comprehensive kidney care strategies
- Policy considerations

### **Bottom-up efforts**

- Patients
- Researchers and innovators

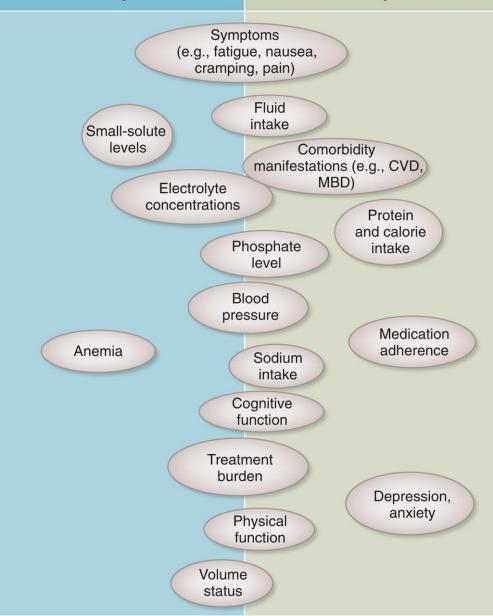


### The goal

- Low-cost options
- Miniaturized devices for greater mobility (wearable, portable, implantable)
- Greener, water-efficient technology
- Technology that more closely mimics kidney function
- Better toxin removal
- Improved mortality and morbidity
- Improvements in physical and mood symptoms
- Robust, safe, complication free

### Directly affected by dialysis treatment or dialysis unit care

### Not directly affected by dialysis treatment or dialysis unit care



### Potential targets for goal-directed dialysis care











### Providing "Adequate" Dialysis and Symptom

Patients may interpret "adequacy" differently than

The term "adequate dialysis" be changed to "goal-directed dialysis,"





KDIGO Controversies Volume 96, Issue 1p37-47July 2019

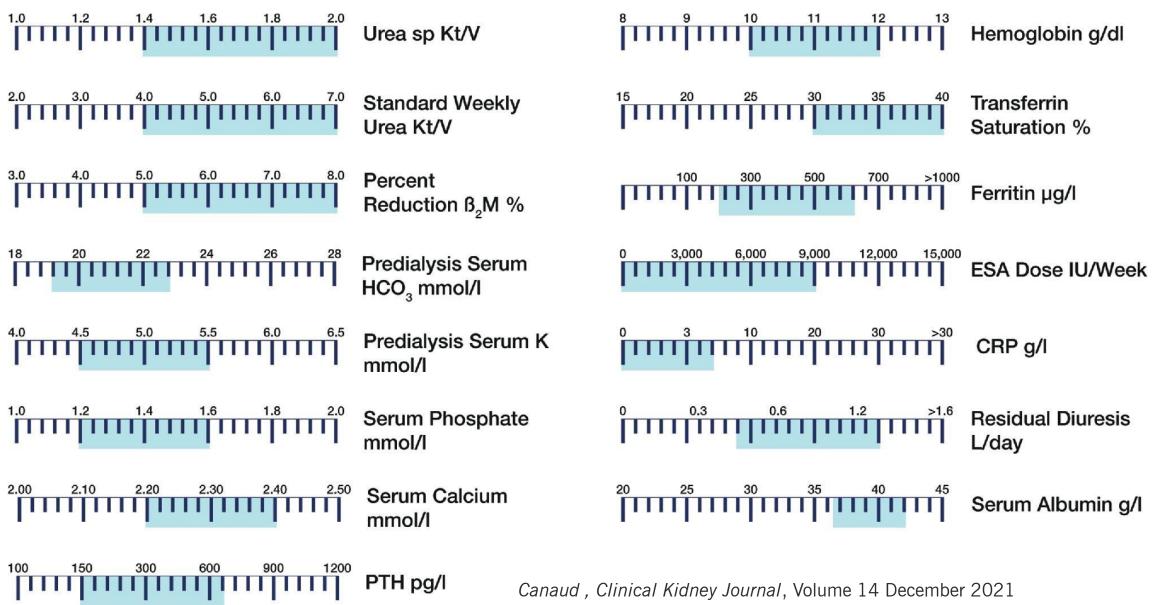
### Changing the HD Adequacy to beyond Kt/V

# Advancing Dialysis

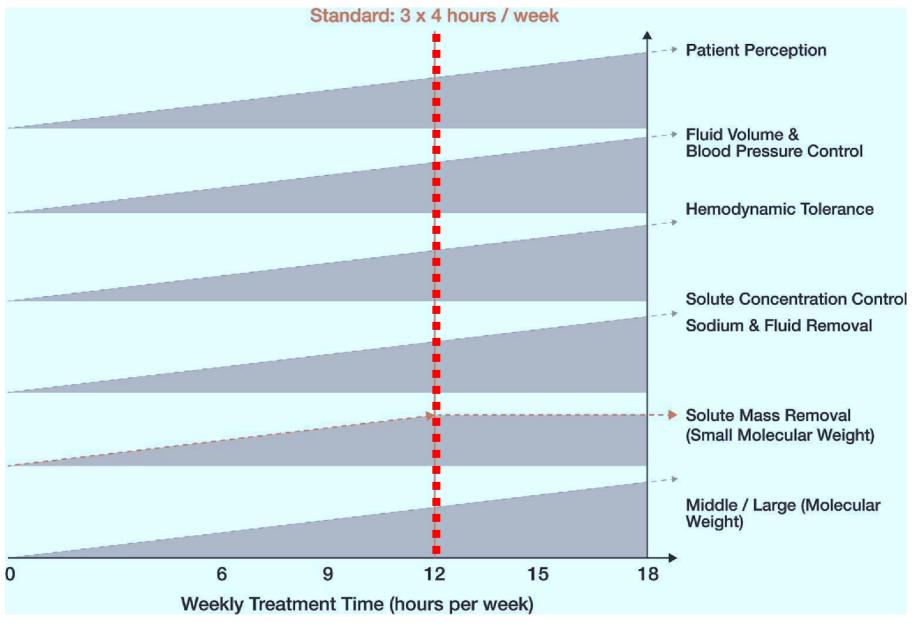
### HD adequacy indicators as a broad and multitargeted approach covering individual patient needs

		Indicator	Target
1.	Patient perception: Lack of symptomatology	Patient feeling Symptom free	
2.	Fluid volume control: Dry weight probing	No edema – No dyspnea BCM – Lung US	
3.	Blood pressure control	Pre – Post dialytic BP Heart Rate	
4.	Hemodynamic stability – Tolerance	Intra Dialytic Hypotension Postdialysis Fatigue – Recovery time	
5.	Dialysis doese monitoring		
	- Small Molecule target	Urea Kt/V (sp/dp) – locnic Kt/V OCM	
	- Middle - Large Molecule target	Standard wk Kt/V \$\beta_2\$-micoroglubulin % reduction / Predialysis	
6.	Acid Base Control - Potassium Control	Serum Bicarbonate pre / post dialysis Serum K pre / post dialysis	
7.	Phosphate – Calcium – Bone Metabolism control	Serum Phosphate pre / post dialysis Serum Calcium pre / post dialysis 25OHD3 – PTH	
8.	Nutritional status control	SGA Albumin – nPCR – Dietary Caloric/Protein Intake	
10	). Inflammation control	Hb - H Iron TSAT - Ferritin	
11	. Preservation of residual kidney function	CRP Diuresis Residual GFR	
12	2. Health related quality of life – Patient reported outcome	HRQOL SF36 EuroQOL	

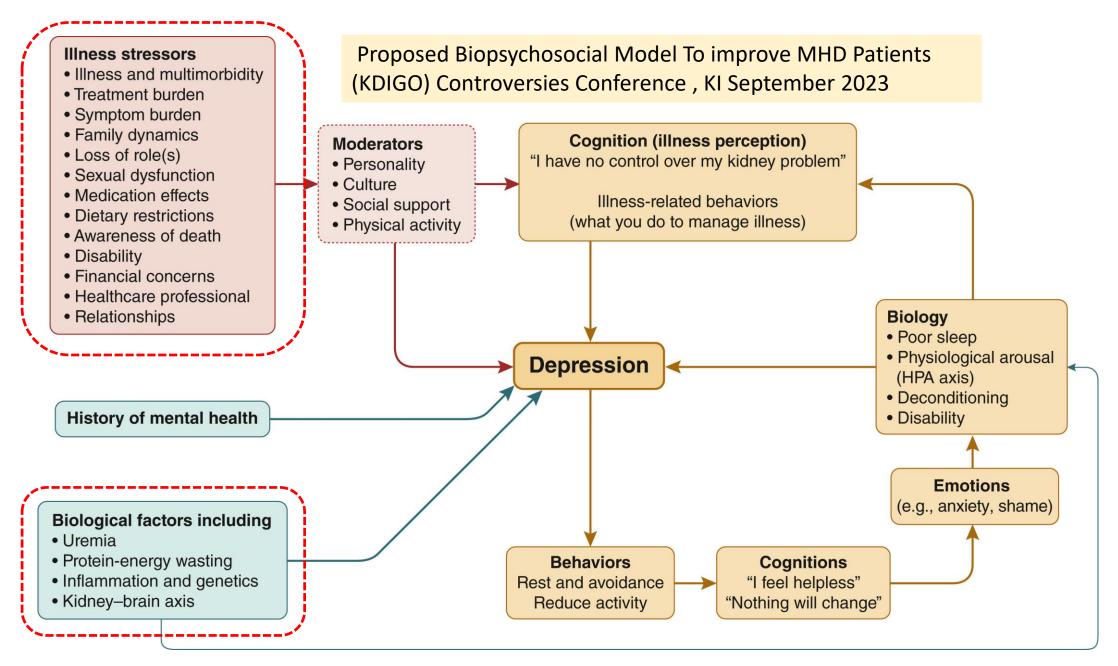
(high on the left end, low on right end)



### Impact of the duration of the therapy session (treatment time)



Canaud, Clinical Kidney Journal, Volume 14 December 2021





### PaCE CKD - The impact of caregiving on health states and work productivity in chronic kidney disease: results from an international survey

Hesham M Elsayed, 1 Helmut Reichel, 2 Janani Rangaswami, 3 Ciro Esposito, 4 Richard Hull, 5 Ricardo Correa-Rotter, 6 Mai-Szu Wu, 7 Steven Chadban, 8 Juan Jose Garcia Sanchez, 9 Surendra Pentakota. 10 Thames Kularatne. 11 Simon Fifer 11

1/Ain Shams University Faculty of Medicine, Cairo, Egypt; 2/Nephrologisches Zentrum Villingen-Schwenningen, Villingen-Schwenningen, Washington DC VA Medical Center; Nephrology Division, George Washington University School of Medicine, Washington, DC; 4/Nephrology, IRCCS Maugeri, University of Pavia, Pavia, Italy; St George's University Hospitals NHS Foundation Trust, London, United Kingdom; National Institute of Medical Sciences and Nutrition Salvador Zubiran, Mexico City, Mexico; Taipei Medical University, Taipei, Taiwan; Royal Prince Alfred Hospital, Camperdown, NSW, Australia; Health Economic and Payer Evidence, BioPharmaceuticals, AstraZeneca, Cambridge, United Kingdom; 10Medical Affairs, BioPharmaceuticals, AstraZeneca, Cambridge, United Kingdom 11CaPPRe: Community and Patient Preference Research, Sydney, NSW, Australia

### **KEY FINDINGS** This multinational survey shows caregiving for patients with chronic kidney disease (CKD) can lead to considerable productivity losses to carers Caregivers of patients with CKD also experience a reduction in health-related quality of life According to CarerQol-7D responses, caregivers report requiring support with their duties, problems with balancing caregiving with their daily life, and financial problems, which contribute to the **substantial** detriment to their quality of CONCLUSIONS This global survey of informal By supporting those living caregivers of those at all with CKD, caregivers stages of CKD establishes experience financial that there is a global quality burden including through of life burden for informal reduced work productivity. particularly when caring for caregivers of those with CKD

#### Introduction Patients with CKD rely on caregivers for support with their disease, particularly as their condition worsens. Informal caregiving is associated with improved adherence to treatment regimens and patient health-related quality of life

- However, the impact of informal caregiving on the HRQoL for caregivers across all stages of CKD is unknown<sup>2</sup>
- Furthermore, limited data are available regarding the global burden of CKD on caregivers' work productivity and financial well-being3

### Objective

(HRQoL)1



To quantify the effect of CKD on informal caregivers' quality of life and work productivity across seven countries compared with matched general populations

#### Methods



Informal caregivers of adults with CKD were enrolled to a non-interventional survey



Recruitment took place in Egypt, Germany, Italy, Mexico, Taiwan, the UK and USA



A general population cohort was also enrolled. matched for age, gender, and area of residence



Work productivity was measured using the Work Productivity and Activity Impairment questionnaire by caregivers and the matched general population



HRQoL was measured by ED-5D-5L index scores versus the matched general population and by dialysis status

EQ-5D-5L could not be assessed in the Italian cohorts

those with advanced disease

CarerQol-7D was measured via several domains including support with carrying out care tasks, financial problems because of care tasks and problems combining care tasks with daily activities Outcomes were stratified by the dialysis status of the patient

### Results

Stage of CKD Stage 1-2

Stage 3-4

Stage 5/KRT

Table 1. Caregiver demographics and duties



26

0 21	30	33			
		33	11	13	12
4 66	51	57	54	57	51
4 8	13	9	35	24	35
2 5	6	1	0	6	2
		14 66 51	4 66 51 57	4 66 51 57 54	14 66 51 57 54 57

Caring for (%)§	for (%)§						
Parent	22	29	25	28	29	44	_
Partner	42	37	40	32	30	26	
Other	36	33	35	41	41	30	



‡ Includes patients self-reporting as CKD stage 5 or dialysis treatment or transplant

Non-dialysis dependent

Dialysis dependent

General population

Non-dialysis dependent

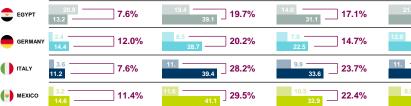
Dialysis dependent

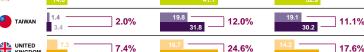
Caregivers

### old tasks, help with personal car UNITED

#### Informal caregivers of patients with chronic kidney disease experience work productivity and activity impairment

Figure 1. Informal caregivers reported worse work productivity and higher impairment scores (mean % time lost) versus the matched general population





Presenteeism



particularly for those caring for patients on dialysis according to CarerQoL-7D measures Figure 2, EQ-5D-5L index scores of informal caregivers of Figure 3. Proportion (%) of informal caregivers of patients with CKD reporting problems

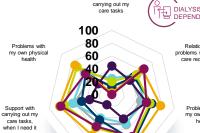
Absenteeism

patients with CKD versus the general population General population -0.20 -0.06 Caregivers Non-dialysis dependent Dialysis dependent General population -0.15 0.00 Caregivers

\_\_-0.13

**III** 1 1



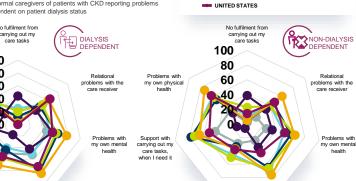


Financial problems

because of my

care tasks

in CarerQoL-7D domains dependent on patient dialysis status No fulfilment from DIALYSIS



Financial problems care tasks

EGYP

= ITALY

TAIWAN

Matched general population

Non-work activity impairment

GERMANY

UNITED KINGDOM

MEXICO

20.5%

39.5%

34.9%

Caregivers

Work impairment

Problems combining my care tasks with my daily activities

my care tasks with my daily activities

Problems combining

1. Ibrahim et al. PLOS ONE. 2015;10(7):e0129015; 2. Adejumo et al. Ghana Med J. 2010;5(43):190-6; 3. Michalopoulos et al Kidney Med 2022 Mar 7:4(4):100439

Dialysis - self reporting as receiving dialysis Non-dialysis - all those who do not report as receiving dialysis This research was supported by AstraZeneca. Medical writing support was provided by Peter Gabb and Rowena Jenkins of Health Economics and Outcomes Research Limited

-0.13



International Trends in Mortality on Hemodialysis Through Changes in Hemodialysis Practices in the Dialysis Outcomes and Practice Patterns Study (DOPPS) *AJKD January 2025*NATIONAL KIDNEY



In Europe, we observed a 13% improvement in overall case-mix adjusted survival per decade. Trends in facility practice measures, especially Kt/V and phosphorus, explained 10% improvement in case-mix survival per decade, (10% explained of 13% improvement) of the observed improvement.



In Japan, 12%/decade improvement in case-mix adjusted survival could be attributed to facility practices, especially Kt/V and IDWG.



In the United States, 47%/decade improvement in case-mix adjusted survival could be attributed to facility practices, especially AVF use and phosphorus control.

FOUNDATION.

### AJKD January 2025

### International Trends in Mortality on Hemodialysis Through Changes in Hemodialysis Practices

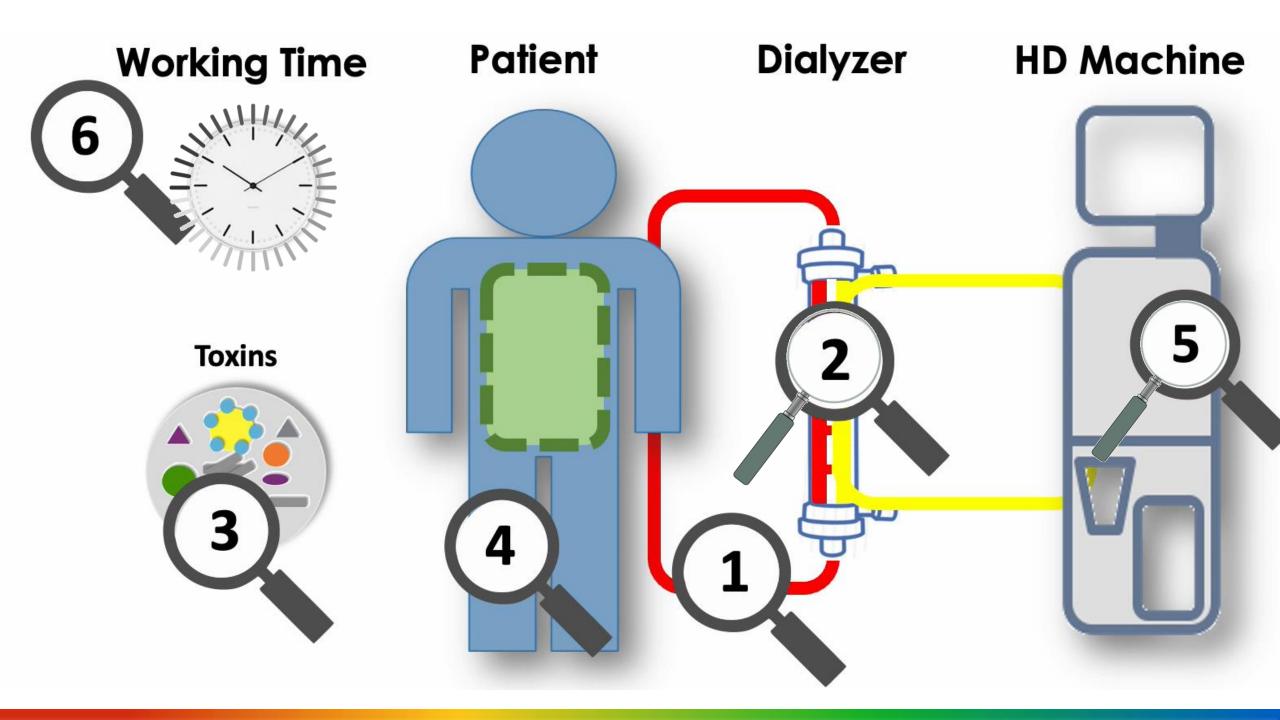
#### **Findings Aim and Population** HD practices change over time, but trends differ by region Proportional improvement in case-mix adjusted survival/decade mediated through practices PRACTICE PATTERNS STUDY 0% -5% 5% 10% 15% 20% 25% 30% 35% 40% 45% Observational prospective Europe cohort study Japan Adult patients treated with hemodialysis: • Europe\*: 26,264 patients US Unexplained improvement • Japan: 27,121 patients • **US**: 34,773 patients ■ Fistula (proportion) ■ Kt/V 1.2+ ■ Phosphorous < 6 mg/dl ■ IDWG < 5.7% \*Germany, Italy, Spain, & UK

**CONCLUSION:** Improvements in adjusted HD patient survival in Europe, Japan, and the US from 1999-2015 can be largely explained by improvements in specific facility practices.

Keith P. McCullough, Hal Morgenstern, Hugh C. Rayner, et al @AJKDonline | DOI: 10.1053/j.ajkd.2024.06.017



### Cognitive impairment Malnutrition and protein-energy Depression wasting Contributors to frailty in • Impaired executive function and Dietary restriction CKD and HD patients neurophysiological activity • Impaired energy utilization Gait imbalance and incoordination **Uremia and inflammation** Sarcopenia Accelerated aging • Impaired anabolic hormone action Immunosenescence Disrupted muscle metabolism Disuse atrophy **Polypharmacy** Drug interactions Adverse drug events • Drug nonadherence **Anemia** Erythropoietin deficiency **Dialysis-related factors** Obesity · Dialysis loss of nutrients and Adipose browning antioxidants • Proinflammatory adipokine production Intradialytic hypotension • Disrupted muscle contraction kinetics Endotoxemia Advanced glycated end-product accumulation CKD-mineral bone disease Bone fragility and muscle incoordination Vascular calcification Impaired response to Gordon, Kidney International, July 2024 erythropoiesis-stimulating agents (ESA) 15



### Could Al improve care for patients with kidney failure?

8th November 2019



### AI in the Dialysis Space

Predicting intradialytic hypotension Improve anemia management Dialysis Adequacy HD Machine K Testing and monitoring





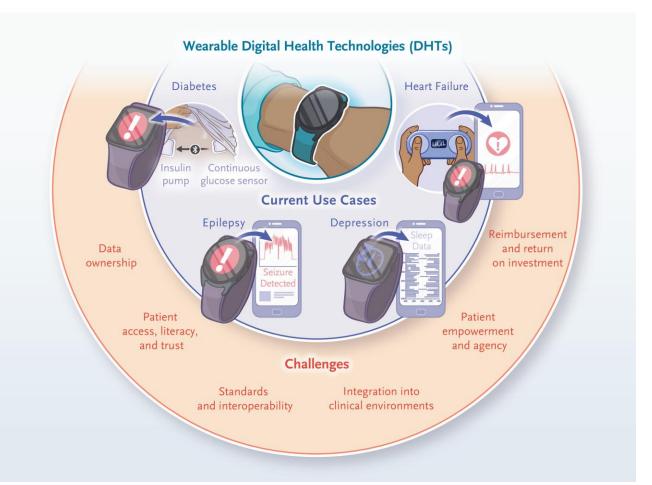


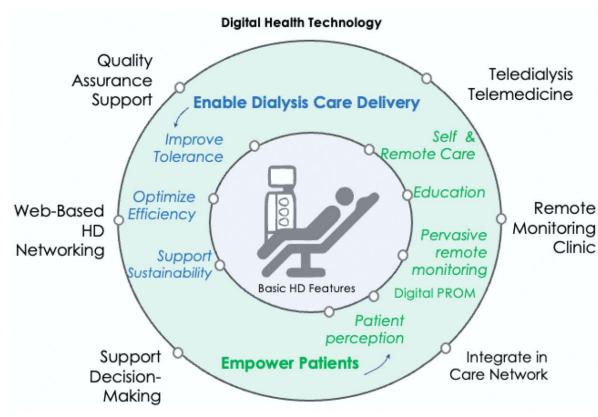
Innovation in dialysis delivery systems is needed to build an adaptive and self-improving process

Minimizing the metabolic and hemodynamic "roller coaster" to change the status quo of dialysis care

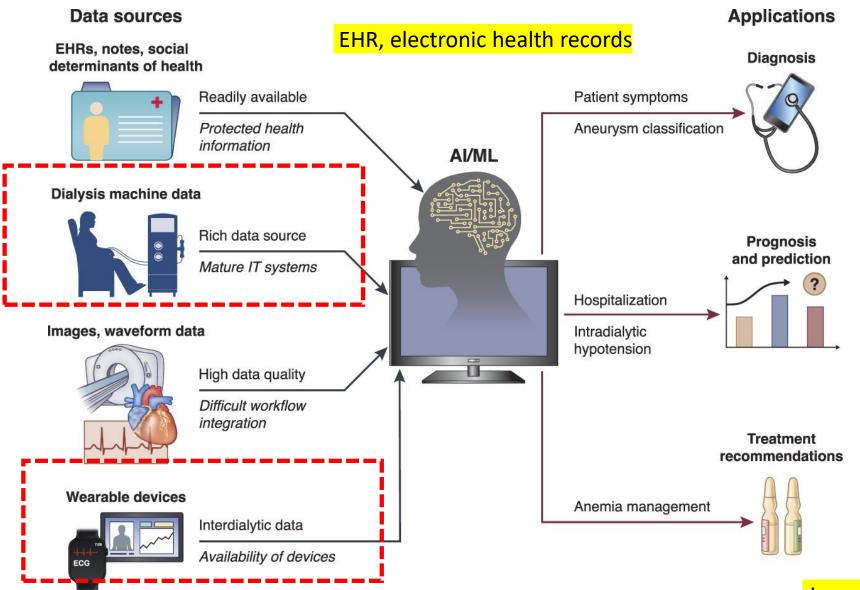
The aim of transforming it from being reactive to being proactive

### Integrating digital health technology (DHT) in HD





Geoffrey, N Engl J Med March 2024



Clinical Journal of the American Society of Nephrology

January 27, 2023.

### Deep Learning Using Electrocardiograms in Patients on Maintenance Dialysis

### **Arrhythmia** prediction

To predict arrhythmia during dialysis based on 10 second EKG at the beginning of dialysis. Allowing real time change in dialysis prescription



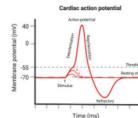
### Electrolyte abnormality prediction







Prediction of electrolytes prior to HD treatment will allow personalized dialysis prescriptions and in the future using wearables or single lead use AI EKG will help predict extreme changes in electrolytes in patients during interdialytic periods







### Structural heart condition prediction

Structural heart abnormalities are common in dialysis population early detection, screening and monitoring with AI-EKG provides a novel opportunity



### **Future directions**

To predict ischemic events, stroke, sudden cardiac death and avoid hemodialysis related cardiac events by personalized real time updated prescription changes

### Intradialytic hypotension prediction

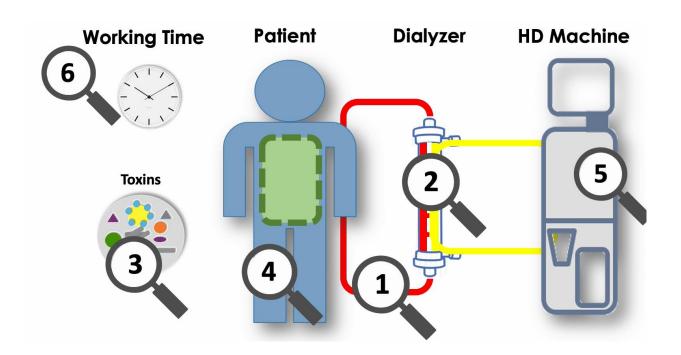
SBP drop ≥20 mmHg or MAP drop ≥20 mmHg

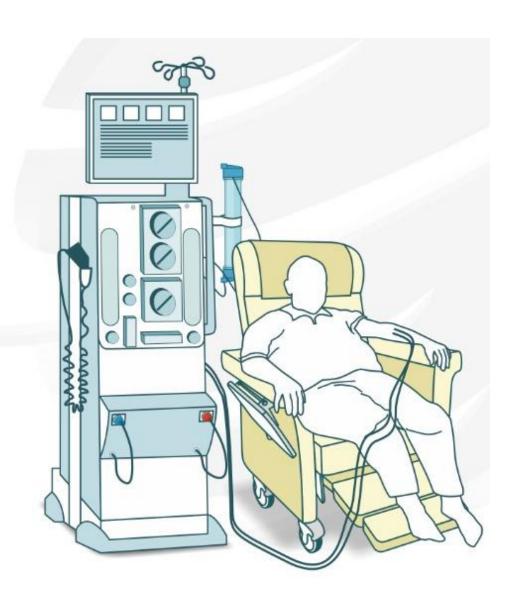
Prediction of IDH based on EKG at the start of dialysis session

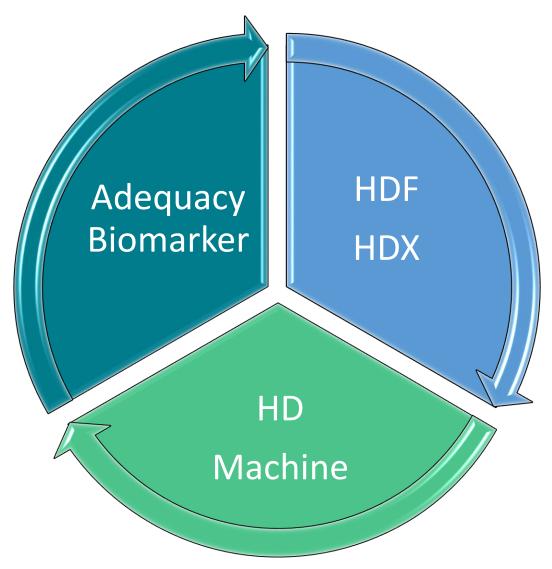




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### Starting hemodialysis The stress factors



Cardia stress Arrythmia



Hypoxemia PH changes



Solute Flux



Inflammation

### THE LANCET

Five trials (n=4153 patients; 2070 receiving HD and 2083 receiving HDF)

This journal Journals Publish Clinical Global health Multimedia Events About

**ARTICLES** · Volume 404, Issue 10464, P1742-1749, November 02, 2024



Haemodiafiltration versus haemodialysis for kidney failure: an individual patient data meta-analysis of randomised controlled trials

Robin W M Vernooij, PhD A Mark Carinna Hockham, PhD b · Prof Giovanni Strippoli, PhD c,d · Prof Suetonia Green, PhD e · Prof Jörgen Hegbrant, PhD f · Prof Andrew Davenport, PhD g · et al. Show more

Affiliations & Notes ✓ Article Info ✓ Linked Articles (1) ✓

HDF can be considered as a superior alternative to the present standard HD

Research Open access Published: 07 January 2025

# Real-world effectiveness of hemodialysis modalities: a retrospective cohort study

<u>Yan Zhang</u> M, <u>Anke Winter, Belén Alejos Ferreras, Paola Carioni, Otto Arkossy, Michael Anger, Robert Kossmann, Len A. Usvyat, Stefano Stuard & Franklin W. Maddux</u>

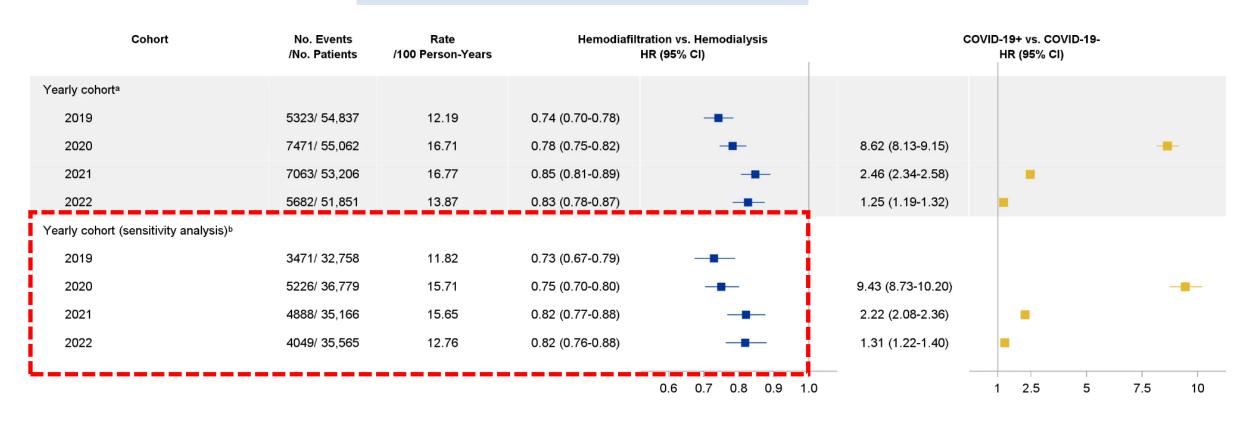
BMC Nephrology 26, Article number: 9 (2025) Cite this article

**756** Accesses Metrics

Retrospective cohort study during 2019–2022

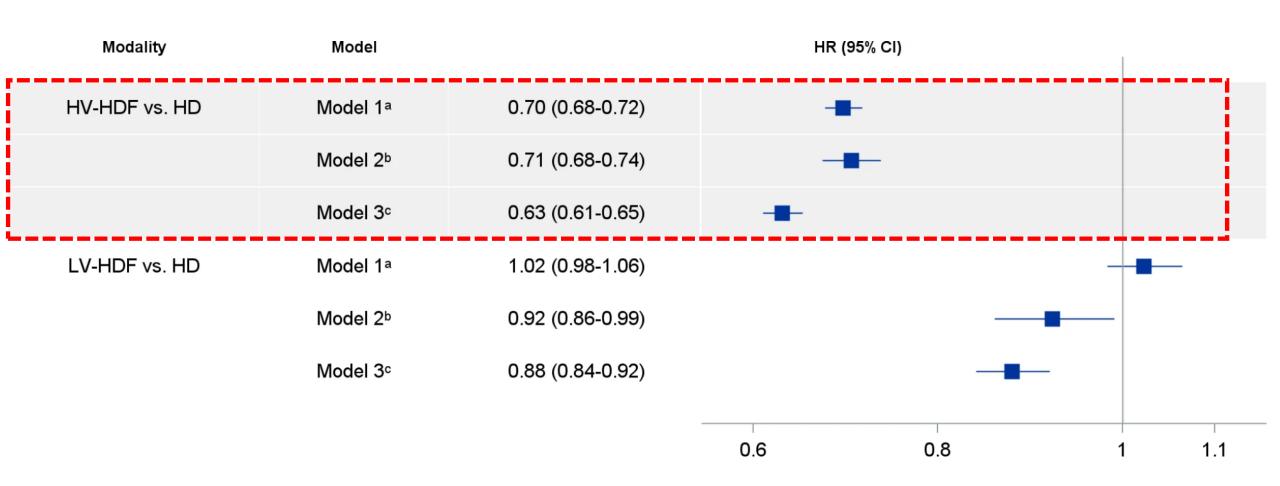
Among patients receiving HDF (mean convection volume ≥ 23 L), the risk of death was reduced by 30% (HR, 0.70 [95% CI, 0.68–0.72]). Hemodiafiltration was also associated with a 31% reduced risk of cardiovascular death.

### All-cause mortality in yearly-cohort analysis.



Zhang, BMC Nephrology volume 26, Article number: 9 (2025)

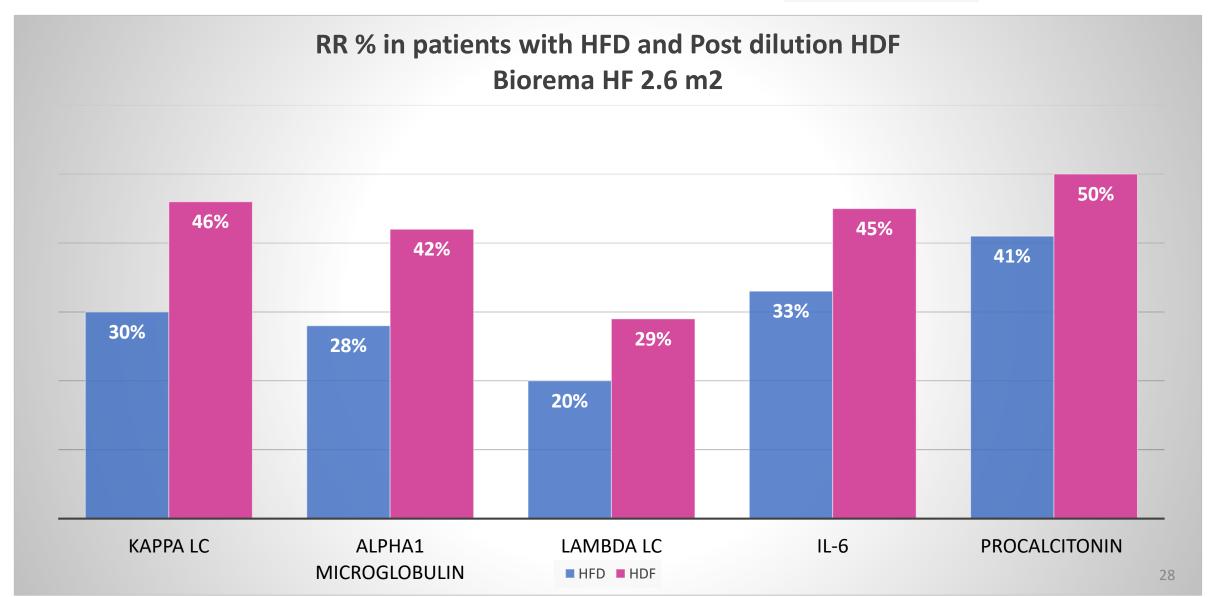
### Association of High Volume HV-HDF and LV-HDF with all-cause mortality relative to HD



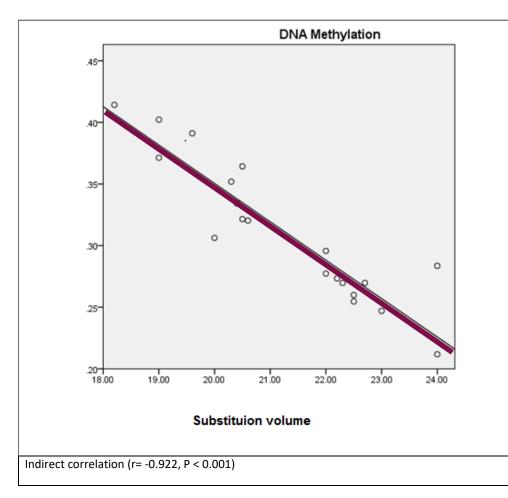
Zhang, BMC Nephrology volume 26, Article number: 9 (2025)

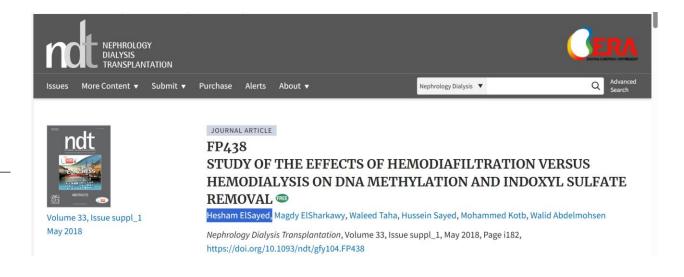
### ASN, Hesham Elayed etail ASN 2022











# EFFECT OF OL-HDF on DNA methylation

Elsayed H, Elsharkawey M etal NDT May 2018

	DNA methylation				
	r	Р			
Substitution volume	-0.922	<0.001**			
**; High Statistical Significant difference					



### EFFECT OF DIALYSIS MODALITY AND MEMBRANE PERMEABILITY ON FGF 23 LEVEL AND CARDIOVASCULAR CALCIFICATION IN ESRD PATIENTS.

Hesham Mohammed Elsayed I, Waleed Ahmed Beshary I, Khaled Mohamed Rezk I, Mostafa Abo Alkhair Mohamed I, Fatma Abdelrahman Ahmed I

## Acute effects of online hemodiafiltration versus high-flux hemodialysis on serum levels of asymmetric dimethylarginine and tumor necrosis factor-alpha

Hesham M. Elsayed, Heba W. El Said, Waleed A. Beshary, Ahmed A. Jaleel, Fatma A. Ahmed

Department of Internal Medicine and Nephrology, Faculty of Medicine, Ain Shams University, Cairo, Egypt

Correspondence to Fatma A. Ahmed, MBBCH, MD, PHD, Ain Shams University, Cairo 11517, Egypt

Tel: +020110121622;

e-mail: fatmagouda@med.asu.edu.eg

Received: 01 June 2022 Revised: 16 July 2022 Accepted: 12 August 2022 Published: 20 January 2023

Journal of The Egyptian Society of Nephrology and Transplantation 2023, 23:3–10

### **Background**

Chronic inflammation as a major determinant of 'dialysis syndrome' is considered as the main factor of morbidity and mortality in dialysis patients. Tumor necrosis factor-alpha (TNF- $\alpha$ ) may play important roles in the development of T helper (Th) imbalance, cardiovascular disease, and wasting in the uremic milieu. Asymmetric dimethylarginine (ADMA) is an endogenous inhibitor of nitric oxide that may be an independent risk factor for endothelial dysfunction and cardiovascular disease. In hemodialysis (HD) patients, plasma ADMA is a strong and independent predictor of overall mortality and cardiovascular outcome.

The aim of this study to evaluate the acute effects of hemodiafiltration (HDF) compared with conventional HD on blood levels of ADMA and TNF- $\alpha$ .

### Patients and methods

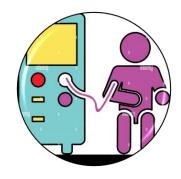
A cross-sectional study was conducted on 20 patients with end-stage renal disease receiving dialysis in the dialysis unit of Ain Shams Specialized Hospital receiving twice weekly HD session with high-flux (HF) dialyzer and once weekly HDF session. Blood samples were collected from all participants before and after HD session and from the same participants before and after HDF session. ADMA and TNF- $\alpha$  levels were assessed by enzyme-linked immunosorbent assay techniques.

### Results

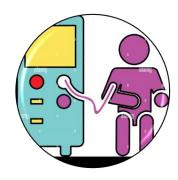
### The optimum we are currently have



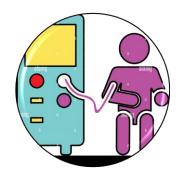
Higher convection therapies



Implementing HDF and Expanded HD



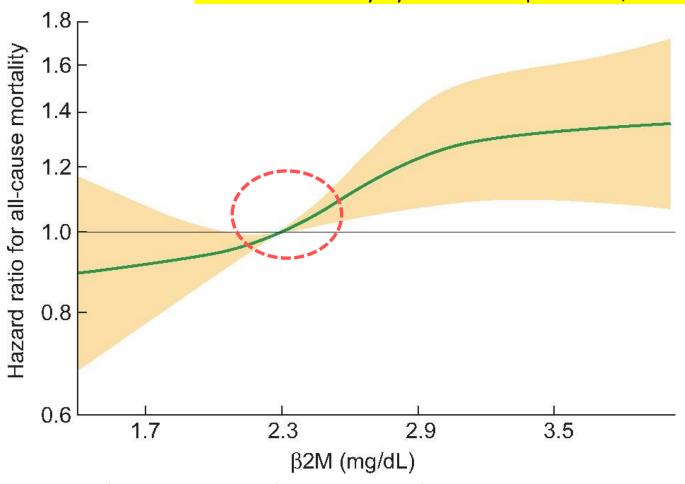
Pre-Dialysis serum B2M



RCT results for ACM and CVM



### All-cause mortality by continuous β2M level, relative to a β2M of 2.3 mg/dL



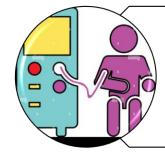
 $\beta$ 2M and mortality in centers routinely measuring  $\beta$ 2M spanned 2011–18 (n = 5332).

Adjusted for age, sex, region (Europe or Japan), DOPPS Phase, dialysis vintage, residual urine volume (≥200 or <200 mL/day), serum albumin and five comorbidities (diabetes, coronary heart disease, congestive heart failure, cerebrovascular disease and other cardiovascular diseases)

Kanda, Clinical Kidney Journal, Volume 14, Issue 5, May 2021



In summary, HDF and HDx represent potential alternatives to conventional high-flux HD with the aim of improving toxin clearance.



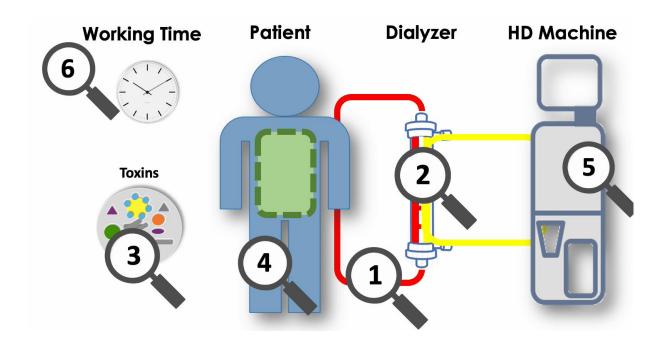
Significant improvements in mortality, cardiovascular outcomes, QoL, and hospitalization with HDF versus conventional HD.



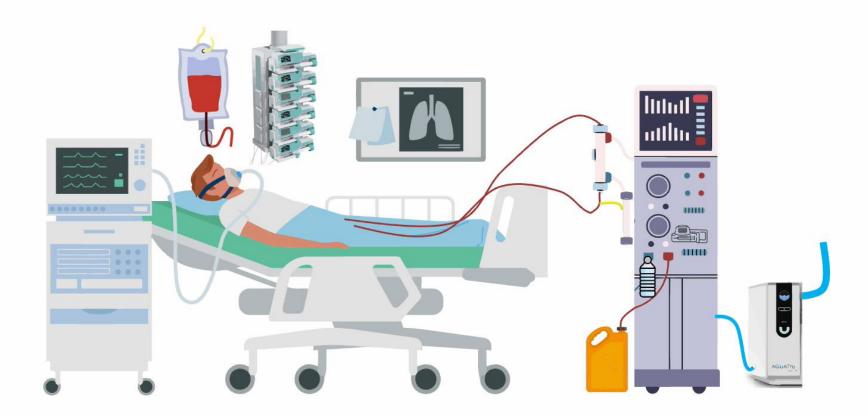
Published studies suggest short-term benefits with HDx in terms of intermediary clinical performance outcomes (e.g., solute clearance); more long-term data on hard clinical endpoints, such as mortality and morbidity, would be beneficial to support clinical decision-making.

### Agenda

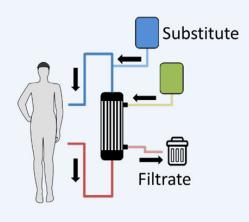
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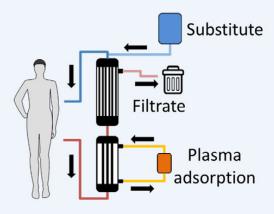
# Typical Environment of AKI Patients in ICU with PIRRT (IHDF) Therapy

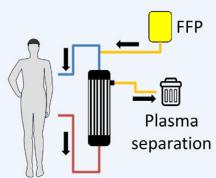


### Extracorporeal Blood Purification Techniques









Convection Therapies

Adsorption Therapies

Combination Therapies

Other Therapies

High Cut-Off
Membranes
(HCO)

Specific Adsorption
Polymyxin B (PMX)
LPS Adsorber

Combined filtration and Adsorption (e.g. oXiris®)

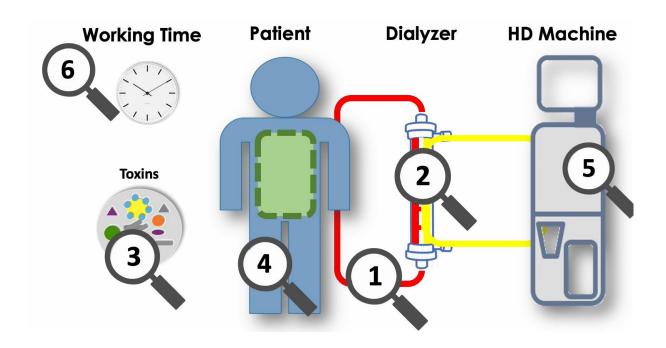
Plasma Exchange

High Volume Hemofiltration (HVHF) Unspecific Adsorption
Hemoadsorption
(e.g. CytoSorb®)

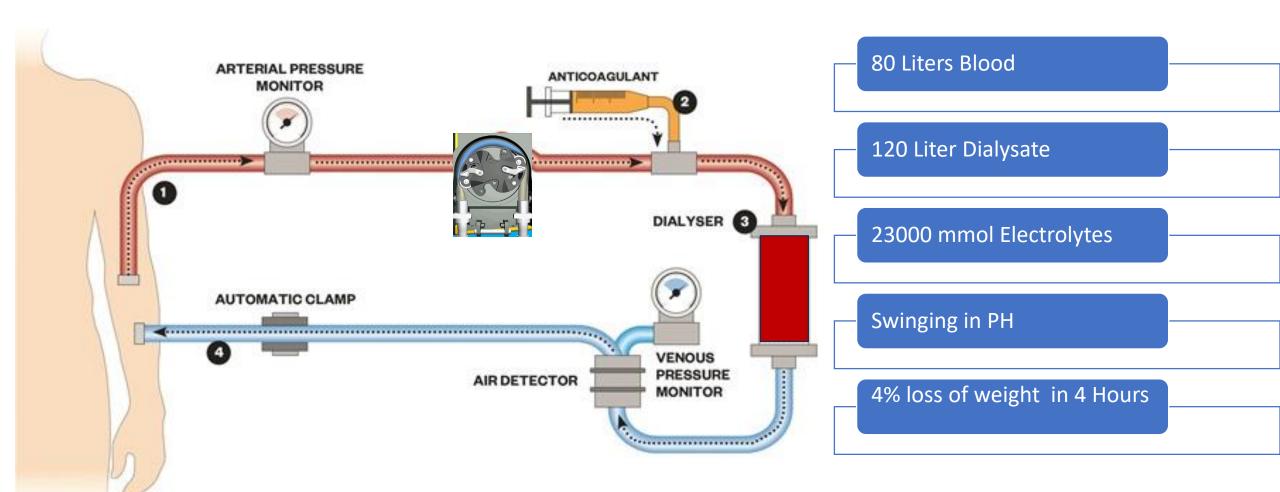
Coupled Plasma
Filtration Adsorption
(CPFA)

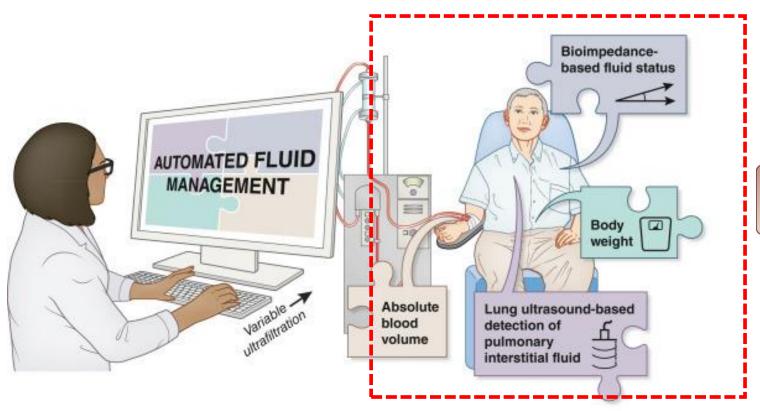
## Agenda

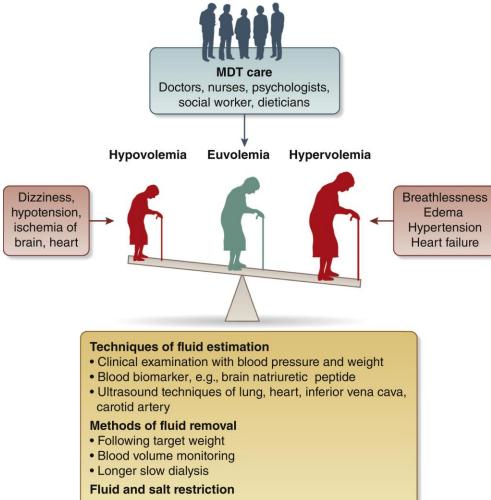
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- 8-Future of intracorporeal HD



#### Sodium First Approach





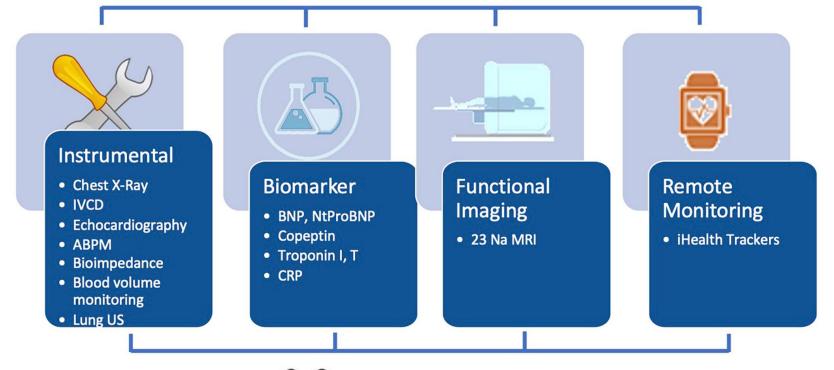




#### Clinical

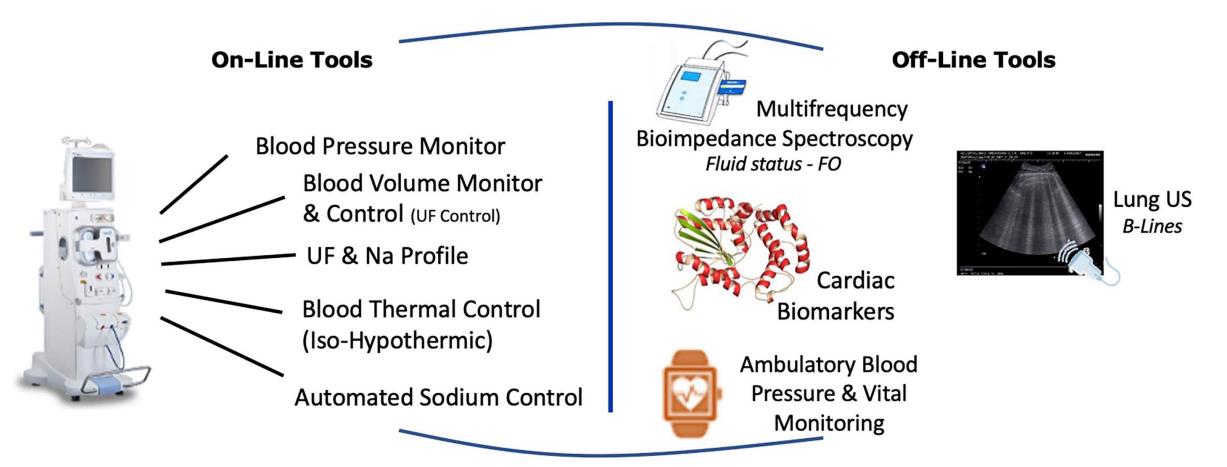
- Clinical Assessment
- Dry Weight Probing
- Blood Pressure
- Weight Loss
- Dialytic Tolerance
- Kidney Function

### Sodium First Approach





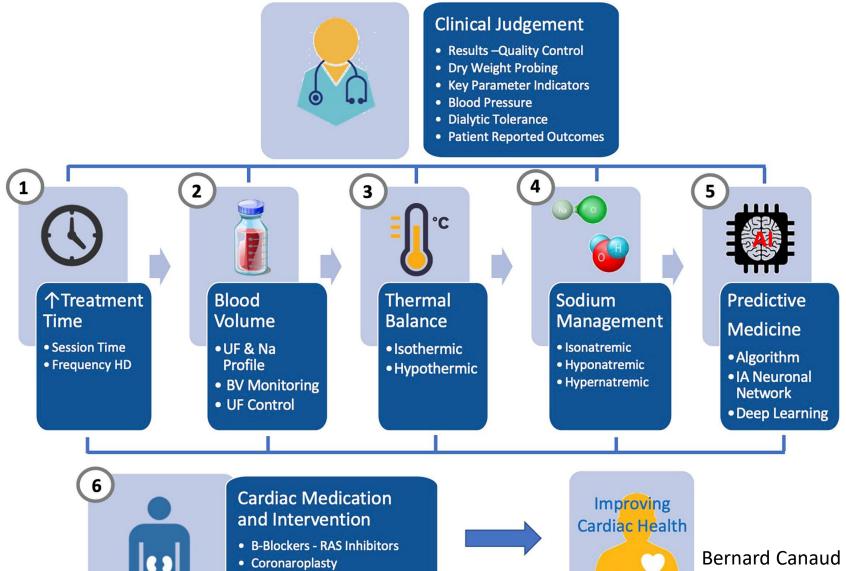
#### **Clinical Decision Support Tools**



**Artificial Intelligence** 

Bernard Canaud Front. Nephrol., 07 July 2022

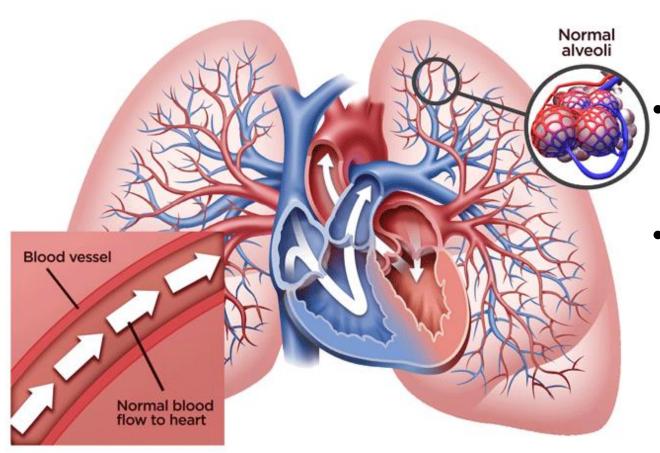
#### Advanced management of sodium, fluid and blood pressure



Cardioversion

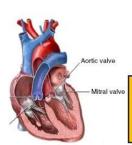
· Pacemaker - Defibrillator

Bernard Canaud Front. Nephrol., 07 July 2022



Pulmonary edema and pulmonary hypertension right ventricular strain

- Pulmonary hypertension during hypoxemia, pulmonary edema can increase right ventricular afterload.
- Consequent right ventricular dysfunction can affect left ventricular preload, leading to hemodynamic instability.

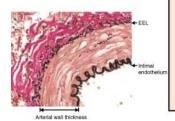


Valvular heart disease Aortic stenosis, mitral regurgitation



#### Coronary artery disease

Medial thickening with smaller luminal area Extensively calcified fibroatheromatous plaques



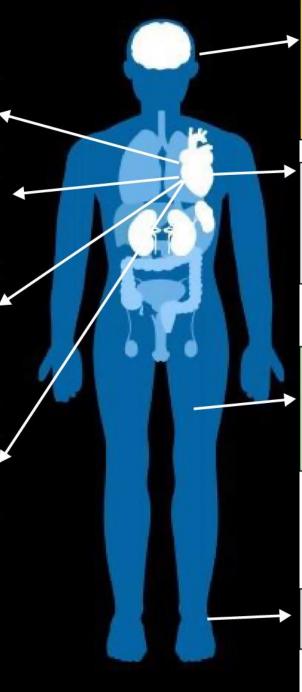
#### Microvascular disease

Extensive arteriolar wall thickening with capillary rarefaction

Ischemia despite normal coronaries and predisposition to vasoconstriction due to endothelial dysfunction

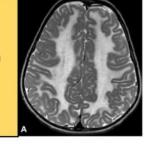


**Dysrhythmias**Atrial fibrillation
Sudden cardiac death



#### Cerebrovascular disease

Ischemic stroke from intimal calcifications
Hemorrhagic stroke from medial calcifications
Small vessel disease including white matter rarefaction
Cerebral microbleeds 5–10 mm in size, and
microinfarcts,

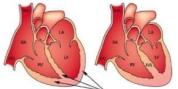


White matter or global atrophy, increased perivascular spaces

#### **Uremic cardiomyopathy**

Left ventricular hypertrophy
Left ventricular systolic and/or diastolic
dysfunction
Subclinical cardiomyopathy with normal left
ventricular ejection fraction





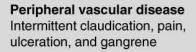
THINNED & WEAKENED WALLS

#### Vascular calcification

Patchy calcification of intima close to atherosclerotic lipid deposits Monkeberg's medial calcific sclerosis with arterial stiffness

Increase in pulse pressure and left ventricular hypertrophy especially seen in diabetics



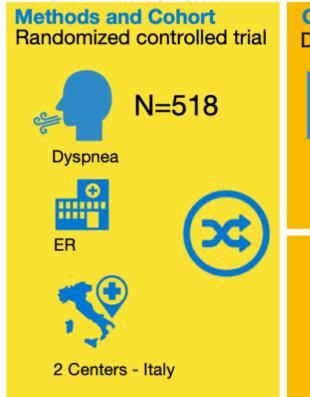




## Does lung ultrasound (LUS) compared to standard approach, improve the accuracy for diagnosis of acute decompensated heart failure (ADHF)?









**Conclusions** Integration of LUS with clinical assessment for the diagnosis of ADHF in the emergency department seems to be more accurate than the current diagnostic approach based on CXR and NT-proBNP.

Reference Pivetta E, et al. Lung ultrasound integrated with clinical assessment for the diagnosis of acute decompensated heart failure in the emergency department: a randomized controlled trial. Eur J Heart Fail. 2019

### Lung Ultrasound to Diagnose Pulmonary Congestion Among Patients on Hemodialysis: Full Versus Abbreviated Scanning Protocols

### Methods

#### **Patient Outcomes**

#### Comparison With 28-Zone Ultrasound







98 patients on maintenance HD evaluated in an ED with a 28-zone lung ultrasound



Median follow up: 778 ± 175 d

B-line score\* < 15: 43% (n = 12)

B-line score\* ≥ 15: 49% (n = 34)

\*on 28-zone study



**AUC (95% CI)** 

A: superomedial	B: inferomedial		
C: superolateral	D: inferolateral		

4 zones



A: 0.83 (0.75-0.9	90) B: 0.83 (0.74-0.90)
C: 0.91 (0.84-0.9	96) D: 0.88 (0.80-0.93)

6 zones



A: 0.86 (0.78-0.93)	B: 0.86 (0.77-0.92)
C: 0.95 (0.88-0.98)	D: 0.91 (0.84-0.96)

8 zones



A: 0.93 (0.86-0.97)	B: 0.88 (0.80-0.94)		
C: 0.94 (0.88-0.98)	D: 0.94 (0.87-0.98)		

**CONCLUSION:** Among patients on maintenance HD presenting to an ED, 4-, 6-, or 8-zone lung ultrasound compared favorably to 28-zone studies for assessing pulmonary congestion.

P = 0.8

Nathaniel Reisinger, Sadichhya Lohani, Jesper Hagemeier, et al (2021)

@AJKDonline | DOI: 10.1053/j.ajkd.2021.04.007



### Heart Failure in HD Patients

Traditional risk factors

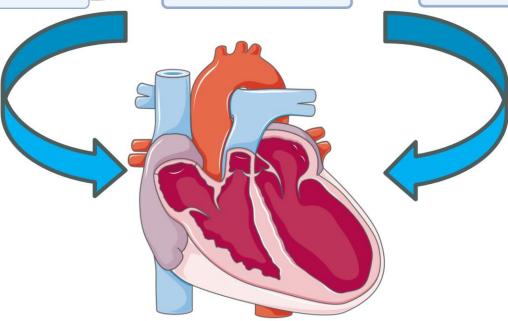
ESRD related risk factors Dialysis related risk factors

- Weight gain
- Hypertension
- Hyperglycemia
- Hyperlipidemia
- Unhealthy lifestyles

- · Capacity overload
- Uremia toxins
- CKD-MBD
- Renal anemia
- Hyperkalemia
- Microinflammation

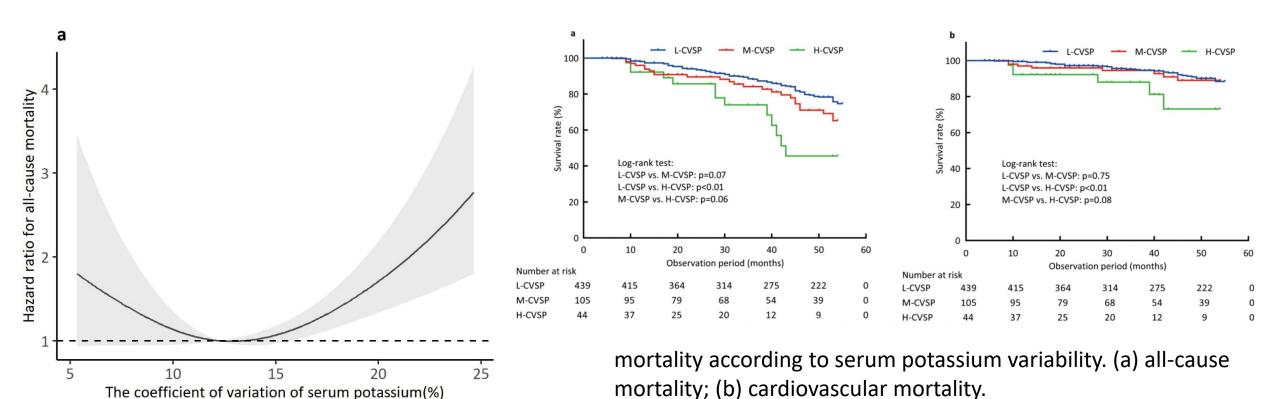
- HD vascular pathway
- HD related complications
- PD related complications

- Reduced vascular compliance
- Myocardial remodeling
- Arrhythmias



- Myocardial ischemia and stunning
- Reduced left ventricular mass
- Myocardial cell damage

# Associations between serum potassium variability and mortality in MHD patients



Hazard ratio of CVSP for all-cause (a) and cardiovascular (b) mortality

588 hemodialysis patients for 45 months

Ru Men, Scientific Reports volume 14, Article number: 29998 (2024)

Effects of dialysate potassium concentration of 3.0 mmol/l with sodium zirconium cyclosilicate on dialysis-free days versus dialysate potassium concentration of 2.0 mmol/l alone on rates of cardiac arrhythmias in hemodialysis patients with hyperkalemia





#### Patients and methods

#### Adults (N = 88) with kidney failure:



Received HD 3 days/week for ≥3 mo with hyperkalemia: two pre-dialysis sK<sup>+</sup> of 5.1–6.5 mEq/l during screening



Had a cardiac loop recorder implanted



Randomized to either a 2.0 K+/2.5 Ca<sup>2+</sup> mEq/l dialysate without SZC or to a 3.0 K+/2.5 Ca<sup>2+</sup> mEq/l dialysate with SZC on non-dialysis days titrated to maintain pre-dialysis sK+ within 4.0–5.5 mEq/l range 8 wks → treatment crossover → 8 wks



Rate of AF events (duration ≥2 min, primary outcome); clinically significant cardiac arrhythmia (CSCA) events; % sK<sup>+</sup> measurements outside 4.0–5.5 mEq/l.

#### **Outcomes**

3.0K+/SZC vs. 2.0K+/noSZC

#### AF episodes ≥2 minutes



Modelled RR (95% CI): **0.52** (0.41–0.65); *P* < 0.001

#### Clinically significant arrhythmias







Modelled RR (95% CI): **0.47** (0.38–0.58); *P* < 0.001

#### sK<sup>+</sup> outside optimal window



OR (95% CI): 0.27 (0.12-0.35)

#### Post-dialysis hypokalemia



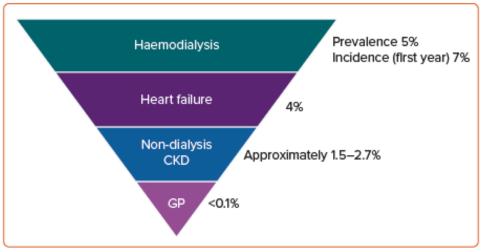
Rate/yr (95% CI): 5.9 (4.5–7.2) vs. 16.5 (14.3–18.7)

Charytan DM, et al. 2024

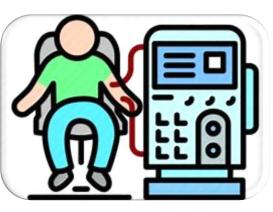
**CONCLUSION** In patients with hyperkalemia on maintenance HD, dK<sup>+</sup> 3.0 mEq/l and SZC on HD days reduced rates of AF, other clinically significant arrhythmias, and post-dialysis hypokalemia compared with dK<sup>+</sup> 2.0K<sup>+</sup>/noSZC<sub>5.0</sub>

# Adjusting the HD prescription against SCD

Figure 2: Annual Rates of Sudden Cardiac Death



CKD = chronic kidney disease; GP = general population. Source: Turakhia et al. 2019. 41 Reproduced with permission from Oxford University Press.



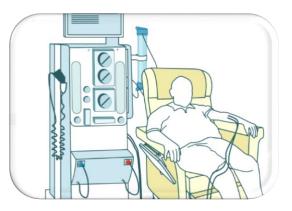
Avoid low potassium Dialysate (< 2mEq/L)



Avoid Low calcium dialysate (<2.5 mEq/L) has been associated with a 40% increase in the risk of SCD.

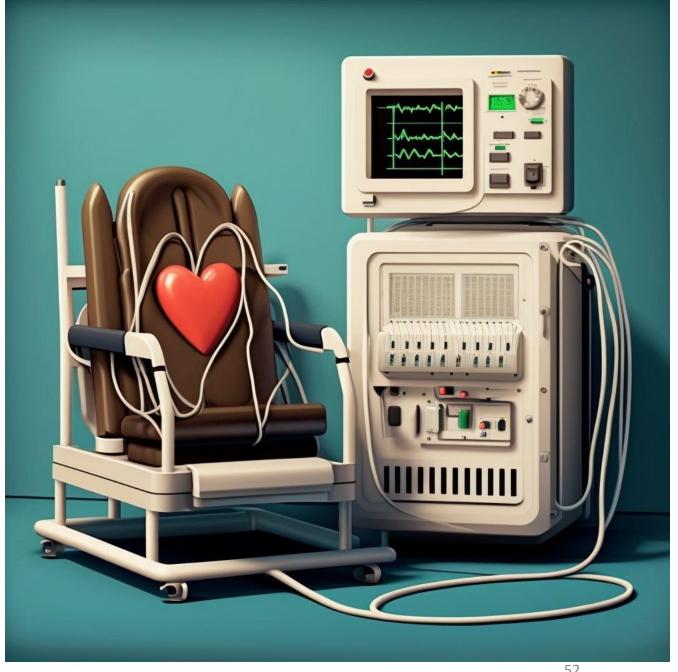


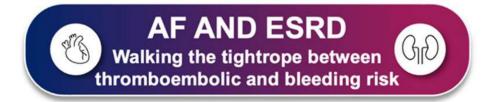
Avoid large shifts in volume from high ultrafiltration rates



Dialysate cooling

## Atrial Fibrillation in MHD patients





## PROTHROMBOTIC STATE

Atrial fibrosis
Ventricular hypertrophy
Accelerated atherosclerosis
Vascular calcification
Endothelial dysfunction
Blood flow abnormalities
Oxidative stress and inflammation
Hypercoagulability

#### PROHAEMORRHAGIC STATE

Anaemia
Reduced platelet activity
Reduced platelet adhesion
Reduced platelet aggregation
Alteration platelet-vessel-wall interaction
Need for antiplatelet treatment
Invasive procedures





Real-world data in this population suggests that apixaban is likely at least comparably effective as warfarin at preventing thromboembolic events and is likely safer when evaluating bleeding risk. Rivaroxaban data is more limited,

Articles Publish Topics About Contact

REVIEW · Volume 10, Issue 1, P40-53, January 2025 · Open Access



## Direct Oral Anticoagulants in Patients With ESRD and Kidney Transplantation

Nicholas W. Lange 🖰 🖾 · Justin Muir · David M. Salerno

Affiliations & Notes ✓ Article Info ✓



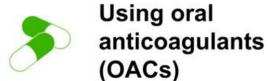
Both apixaban and rivaroxaban display alterations in their pharmacokinetic profiles with renal insufficiency, though neither are appreciably removed by hemodialysis

Drug	Standard Dosing	Dose adjustments	Elimination	Protein binding	Half-life
apixaban <sup>7</sup>	AF: 5 mg twice daily VTE: 10 mg twice daily × 7 days, then 5 mg twice daily	AF: adjust to 2.5 mg twice daily if at least 2 factors: age ≥ 80 yrs, weight ≤ 60 kg, creatinine ≥ 1.5 mg/dl	27% renal clearance (unchanged drug) Biliary and intestinal excretion in feces	87%	12 h
rivaroxaban <sup>8</sup>	AF: 20 mg daily VTE: 15 mg twice daily × 21 days, then 20 mg daily	AF: 15 mg daily for CrCl ≤ 50 ml/min VTE: avoid use for CrCl < 15 ml/min	36% renal clearance (unchanged drug) Excretion in feces	92%–95%	5–9 h

### Oral Anticoagulant Agents in Patients With Atrial Fibrillation and CKD: A Systematic Review and Pairwise Network Meta-analysis

## **Setting & Participants** 8 RCTs and 46 observational studies







170,059 participants

#### Methods

1. Pairwise meta-analysis

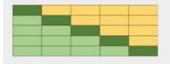


**DOACs (direct OACs)** vs Warfarin



**OACs vs No OACs** 

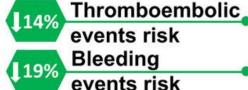
2. Bayesian network meta-analysis

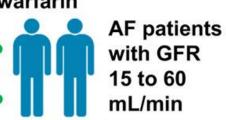


DOAC vs DOAC



Results





- OACs without significant benefits





AF patients

 Dose-adjusted apixaban seem to be superior to other DOACs



AF patients with GFR 25/30 to 60 mL/min

**CONCLUSION:** In patients with AF and mild to moderate kidney disease, DOACs are superior to warfarin, while the preferred agent of DOACs still cannot be determined.

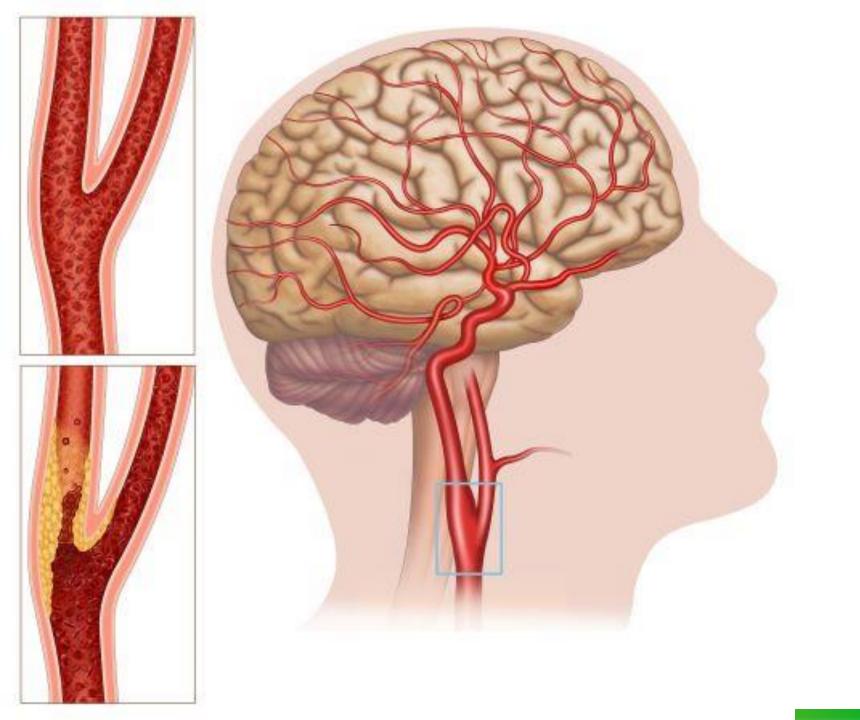


## Comparative Safety and Effectiveness of Warfarin or Rivaroxaban Versus Apixaban in Patients With Advanced CKD and Atrial Fibrillation

Setting & Participants	Findings			
Propensity score matched cohort study			Warfarin vs Apixaban	Rivaroxaban vs Apixaban
2 nationwide US claims			HR (95% CI)	HR (95% CI)
<b>L</b> databases, 2013-2022		Major bleeding	1.85 (1.59-2.15)	<b>1.69</b> (1.33-2.15)
Patients with atrial fibrillation (AF) and CKD stages 4-5 newly initiated on:		Ischemic stroke	<b>1.14</b> (0.83-1.57)	<b>0.71</b> (0.40-1.24)
Warfarin Rivaroxaban vs apixaban (N = 12,488) (N = 5,720)	=	All-cause mortality	1.08 (0.98-1.18)	<b>0.94</b> (0.81-1.10)
CONCLUSION: In patients with AF and advanced CKD, rivaroxaban and warfarin				

**CONCLUSION:** In patients with AF and advanced CKD, rivaroxaban and warfaring were associated with a higher rate of major bleeding compared with apixaban.





Cerebrovascular disorders in HD

## Cerebrovascular Stress: increase the risk of stroke; dialysis patients experience a 10-fold higher incidence, with case fatality rates reaching 90%



High hemodynamic fluctuations



Higher incidence of AF



Up to 10-fold increased risk of stroke in dialysis patients



Vascular calcification (VC)

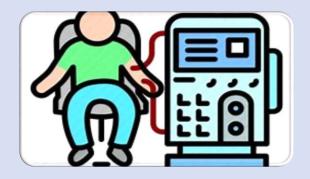


DM HTN HD Vintage

### Seizures caused by electrolyte and glycemic disturbances









#### Hyponatremia

Rapid decreases in serum sodium concentrations can trigger generalized tonic—clonic seizures

Hypocalcemia

increased neuronal excitability due to reduced extracellular concentration of calcium

Hypomagnesemia usually at levels <1 mEq/L

more pronounced in hemodialysis with low glucose dialysate.
Nonketotic hyperglycemia most commonly occurs in older diabetic adults and can cause focal motor seizures.

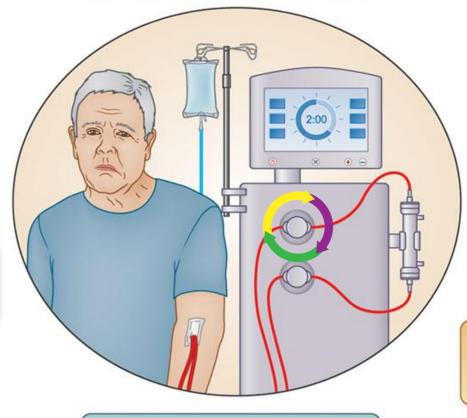
Consider delaying dialysis if patient particularly at risk of raised ICP

Consider continuous forms of therapy (e.g. CVVHD) if patient particularly at risk of raised ICP

For peritoneal dialysis patients, try to minimize hypertonic large volume glucose exchanges

Consider shortening the dialysis session

Time dialysis sessions with essential rehabilitation services



Gentle fluid removal. Excessive ultrafiltration may reduce systemic arterial blood pressure and increase cerebral ischemia

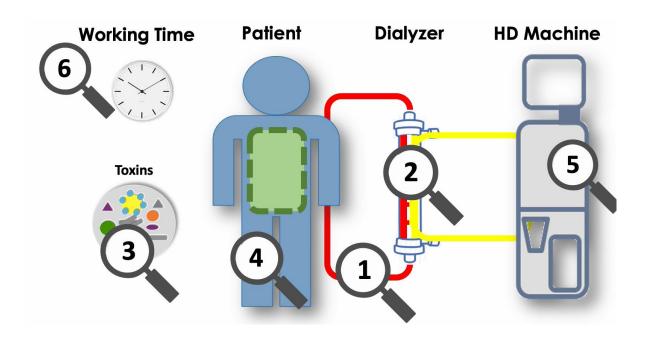
Consider using cooled dialysate

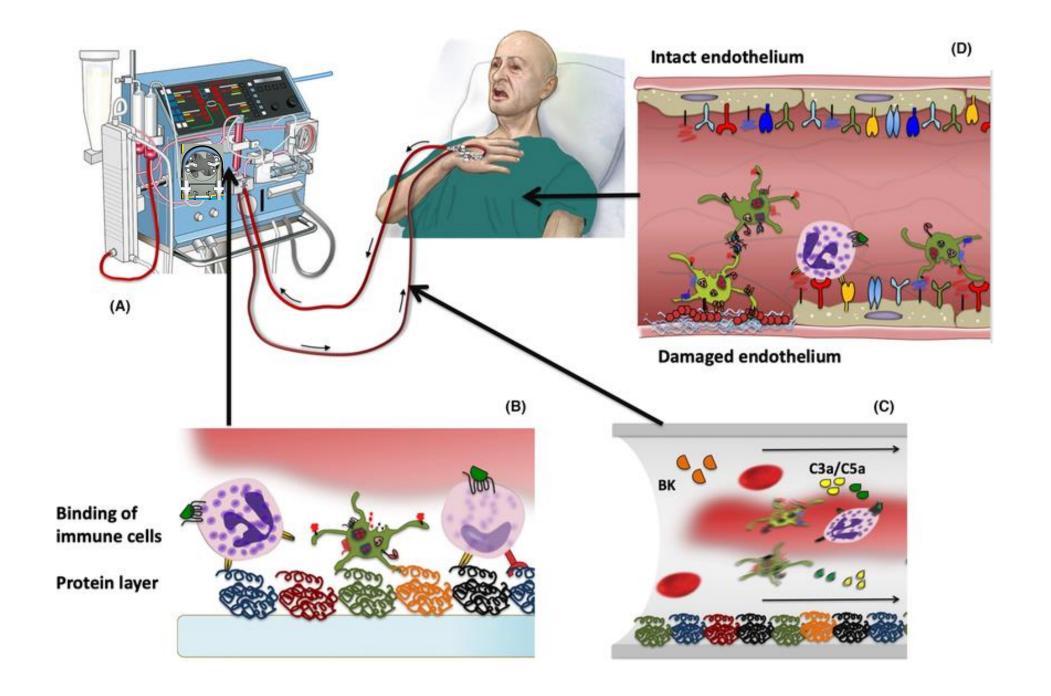
Exercise caution with anticoagulation in acute ischemic stroke and avoid in ICH

Avoid using dialyzers with large surface areas

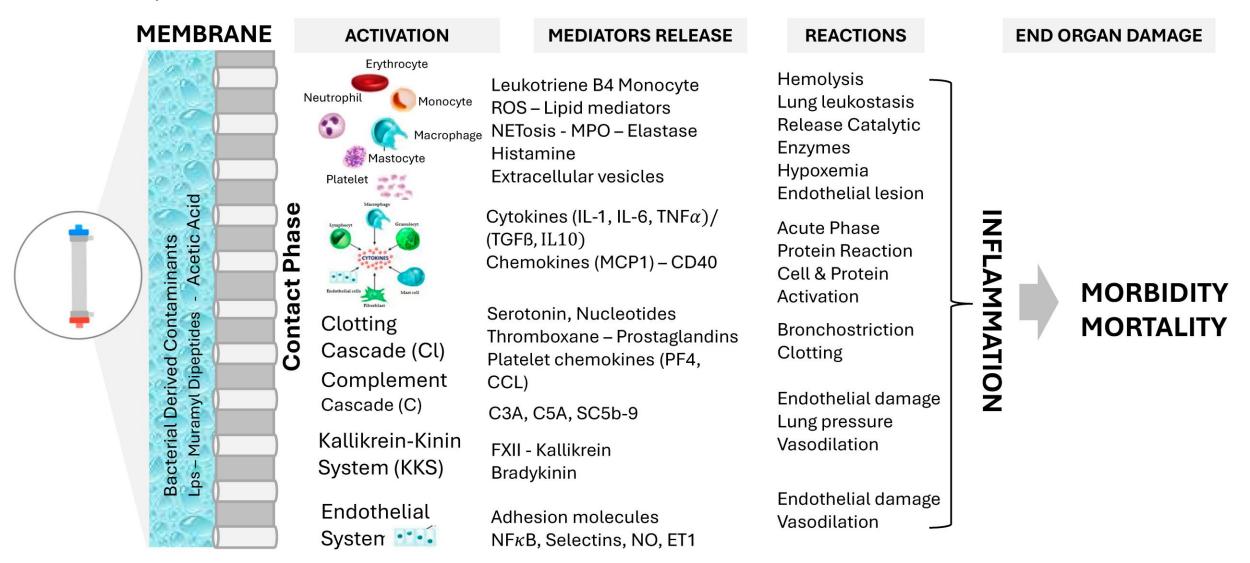
Avoid large hemodynamic shifts.
Start blood flow slowly and increase gradually

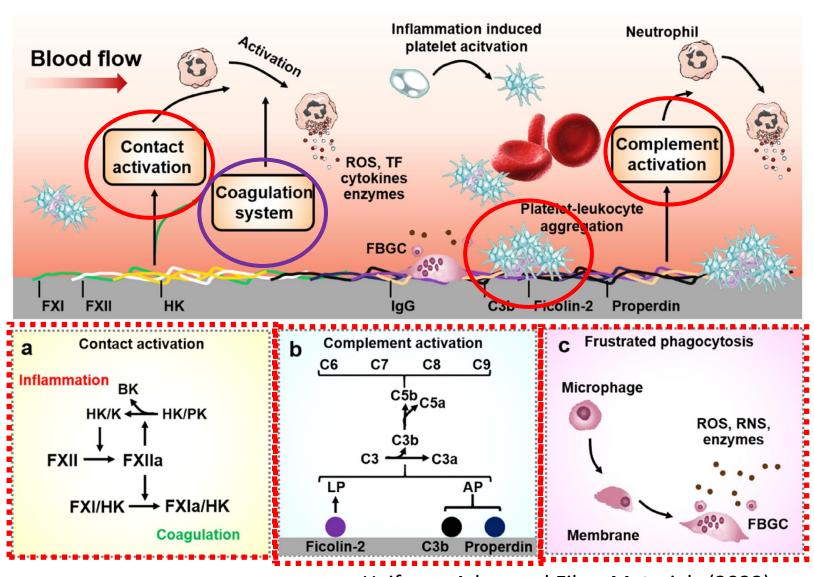
- 1- Patient Centered Approach
- 2- Convection therapies , HDF and HDX
- 3- Extracorporeal therapies in ICU
- 4- Volume and Electrolyte control
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- 6- Cellular activation during HD
- 7- Home Hemodialysis
- 8-Future of intracorporeal HD





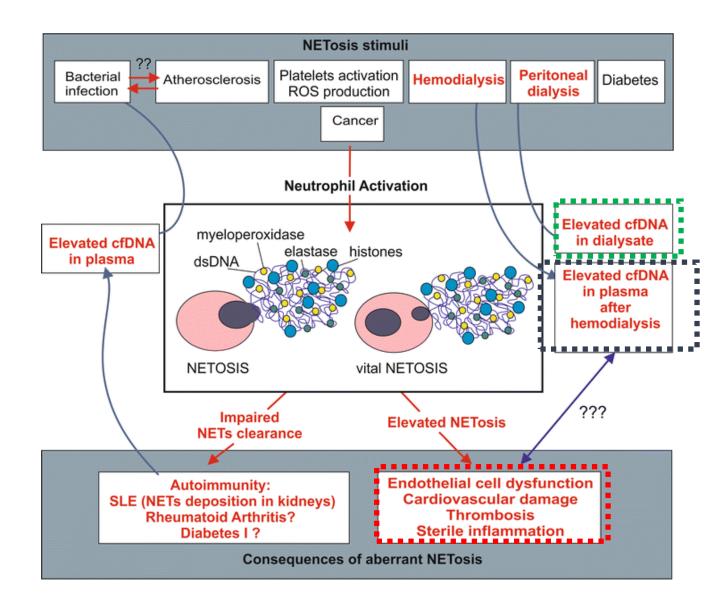
Hemoincompatibility reactions induced by membrane contact and dialysis fluid contaminants result in the activation of various protein cascades and cells





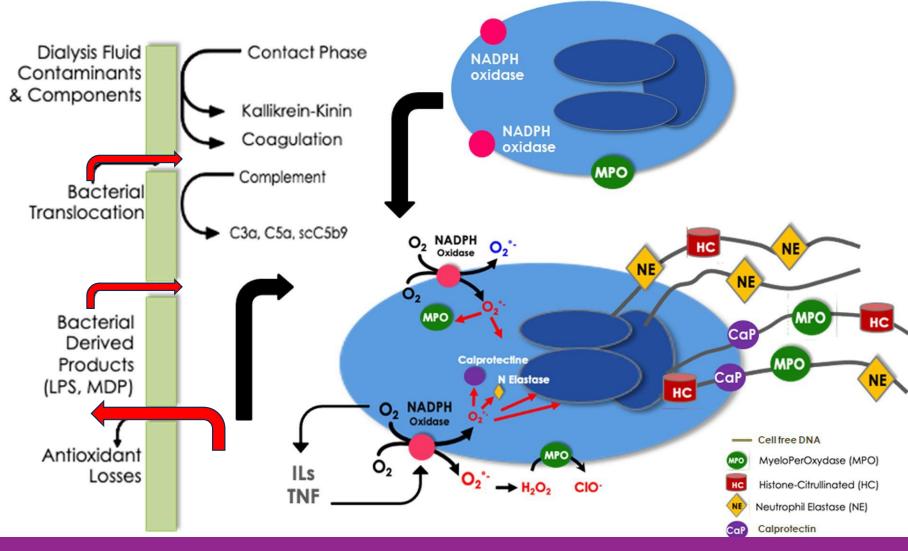
Haifeng, Advanced Fiber Materials (2023)

HD induced inflammation.



Elevated plasma levels of cell-free DNA after a hemodialysis procedure may be activated by NETosis which occurs as a consequence of activation of neutrophils during the process of hemodialysis.

Plasma cell-free DNA levels



Neutrophil extracellular traps in bioincompatibility: formation and factors implicated in dialysis-induced systemic stress. Crestol, Fronters,

Volume 10 - 2023

DOI: 10.1111/1744-9987.13975

#### ORIGINAL ARTICLE



# Effect of dialyzer geometry on coagulation activation in the extracorporeal circuit in maintenance hemodialysis patients: Prospective randomized trial

**Hesham ElSayed** | Khalid Samir Sayed | Mohamed Sary Gharib ©

Division of Nephrology, Department of Internal Medicine, Faculty of Medicine, Ain Shams University, Cairo, Egypt

#### Correspondence

Mohamed Sarv Gharib, Division of

#### Abstract

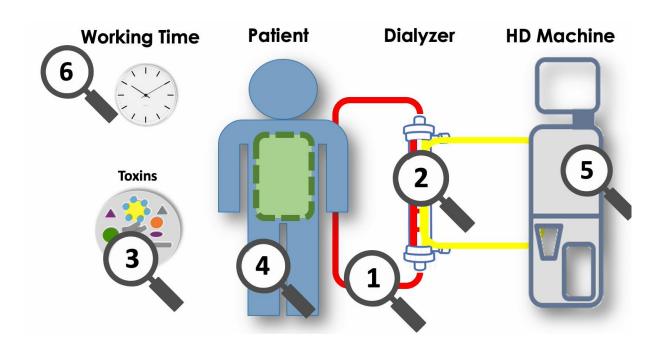
**Background and Objectives:** The coagulation cascade is activated during hemodialysis (HD) due to interaction of blood with the dialysis circuit. There is a paucity of data on the effect of the physical structure of the dialyzers on coagula-

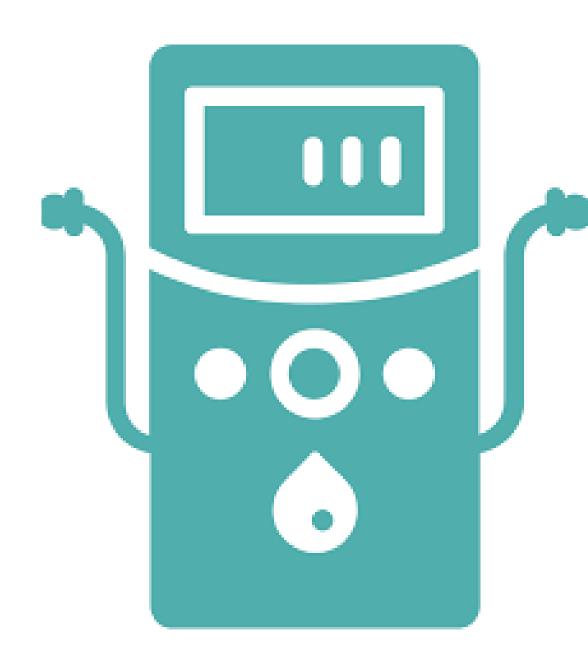
#### Comparison of coagulation activation parameters according to dialyzer membrane type

Parameter	Helixone FX80	Platinum H4	z/t value	P value
TAT at T0h (μg/l), median (IQR)	1.90 (0.52, 2.57)	1.80 (0.82, 2.50)	-0.637	0.524
TAT at T4h (μg/l), median (IQR)	3.00 (2.02, 3.77) <sup>a</sup>	2.65 (1.92, 3.97) <sup>b</sup>	-0.583	0.560
TAT absolute change, (μg/l), median (IQR)	1.15 (0.65, 1.75)	1.15 (0.67, 2.05)	-0.894	0.371

## Agenda

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## Portable Home Hemodialysis

#### **Standard Duration**

**Thrice Weekly In-Center Hemodialysis** 



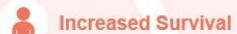
VS

418 ± 54 min



**Extended Duration**Thrice-Weekly Home Hemodialysis





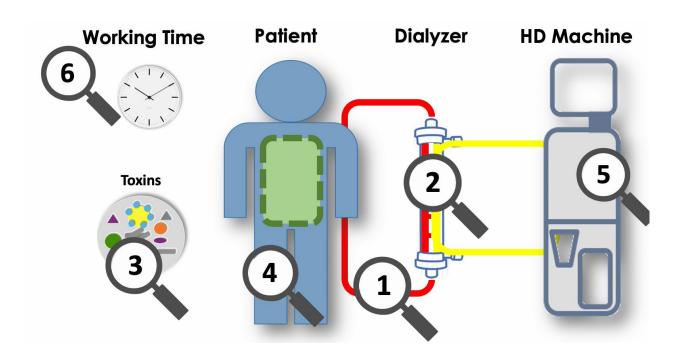


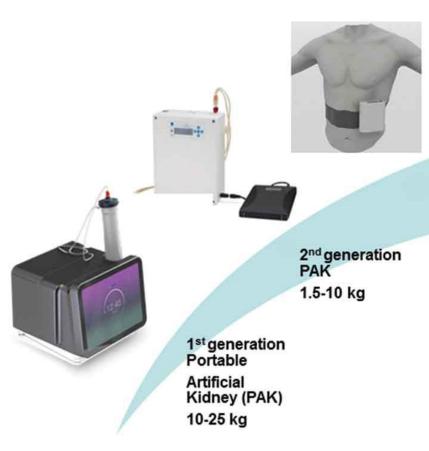


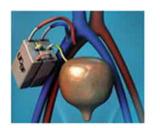


# Agenda

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Wearable Artificial Kidney (WAK) <1.5 kg (Implantable) Bioartificial Kidney (BAK) The road map of dialysis improvements



Creating a bioartificial kidney as a permanent solution to kidney failure

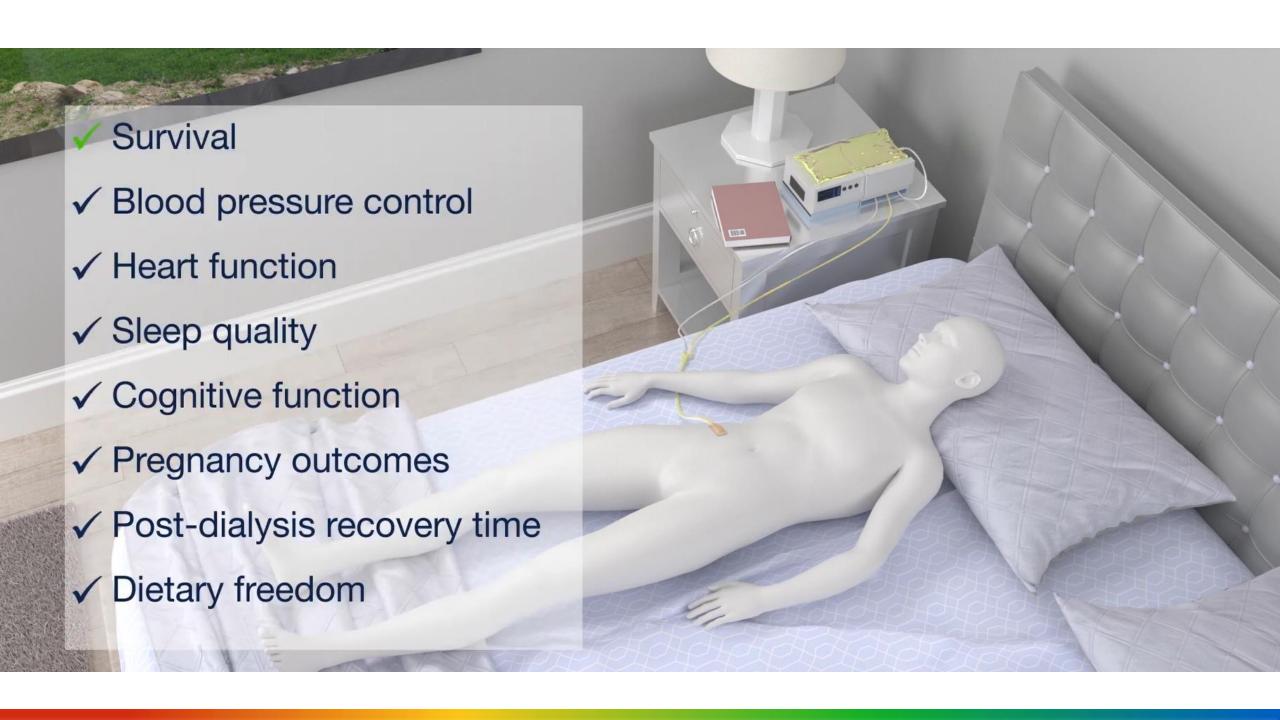
The cells in the bioreactor are isolated from the patient's immune system by the scaffold

Anticipate a GFR value of 20-30 ml/min with this device implanted.

## IHEMO, Hemofilter and Portable dialysate pump









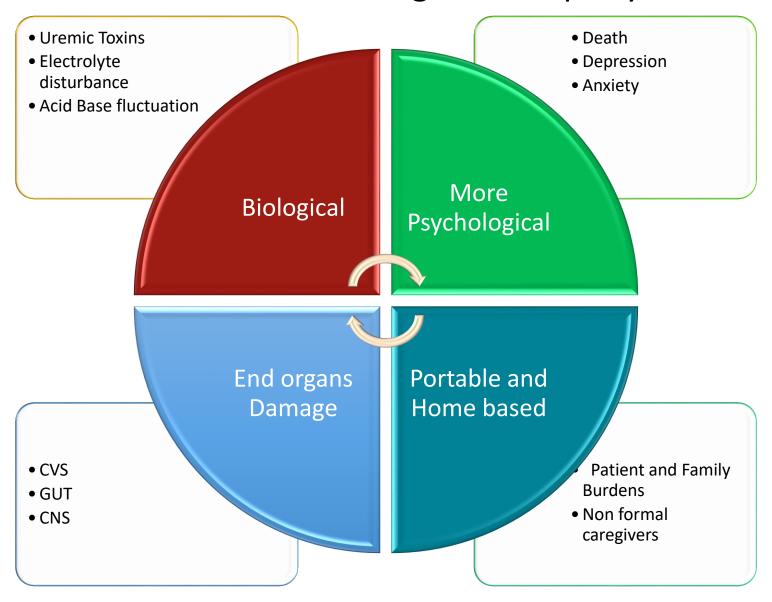


Always there is a lot to think but actually a little we can have

Innovation is the bridge between Ideas and their reality

My bilevels hesham Elsayed

### **Conclusion Defining HD Adequacy**







ACADEMIC DIRECTOR
HESHAM EL SAYED
Professor of Nephrology.





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Emeritus Professor of Nephrology,
Montpellier University, France.



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