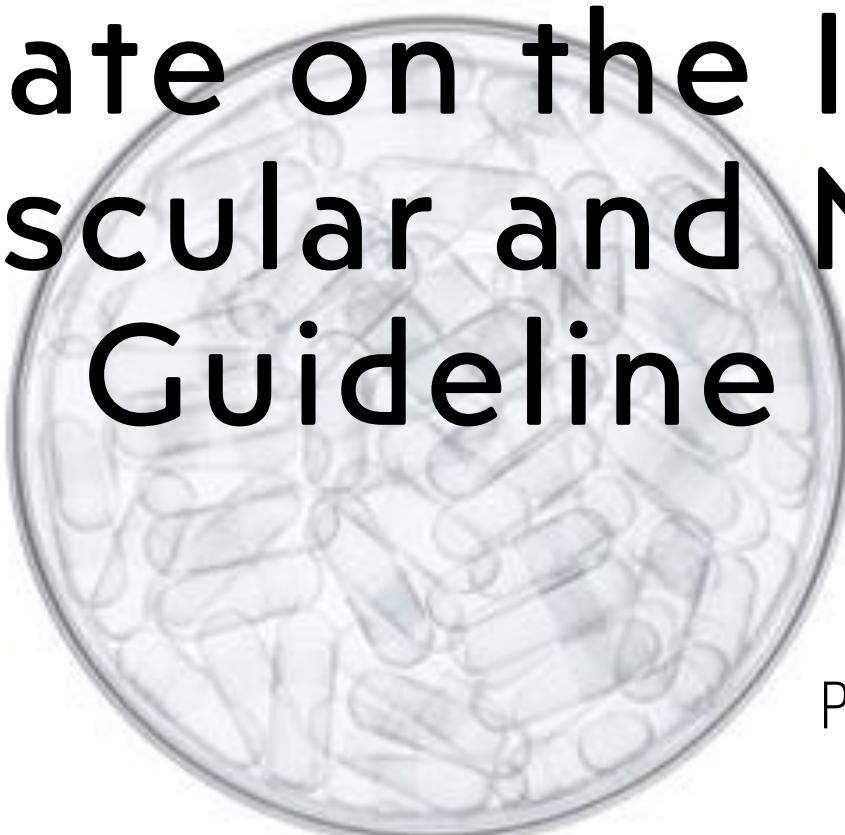


Update on the ISPD Cardiovascular and Metabolic Guideline



Dr. KM Chow

Prince of Wales Hospital, Chinese
University of Hong Kong



DECLARATION

- No conflict of interest
- This is Draft of the ISPD Guidelines, not yet published



OUTLINE

- How we prepared
ISPD Guidelines
- What's New in ISPD
Guidelines



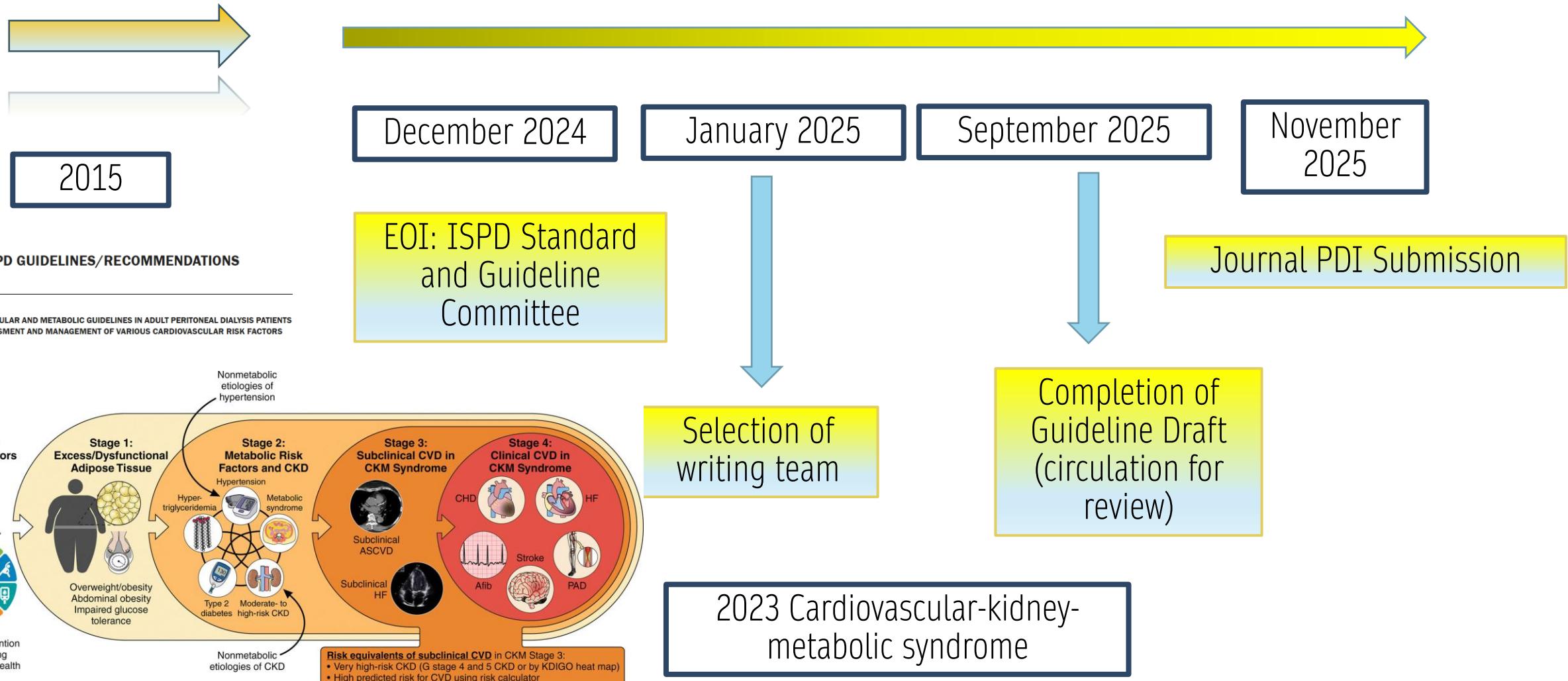
ISPD GUIDELINES/RECOMMENDATIONS

ISPD CARDIOVASCULAR AND METABOLIC GUIDELINES IN ADULT PERITONEAL DIALYSIS PATIENTS PART I – ASSESSMENT AND MANAGEMENT OF VARIOUS CARDIOVASCULAR RISK FACTORS

Angela Yee Moon Wang,¹ K. Scott Brimble,² Gillian Brunier,³ Stephen G. Holt,⁴ Vivekanand Jha,⁵ David W. Johnson,^{6,7} Shin-Wook Kang,⁸ Jeroen P. Kooman,⁹ Mark Lambie,¹⁰ Chris McIntyre,¹¹ Rajnish Mehrotra,¹² and Roberto Pecoits-Filho¹³

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To Be Updated after 10 years



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13. Dr Wilson Wai Hung Tang (Cardiologist, USA)

Composition and Diversity

- 9 Nephrologists
- 2 Nurses
- 1 Dietitian
- 1 Cardiologist
- Male: female ratio 6:7
- LMIC author 8%

GRADE

Grading of Recommendations Assessment, Development
and Evaluation

GRADE

Certainty ratings	General principles of rating
D - Very low	Opinion-based
C - Low	Observational or registry studies Randomized controlled trials with serious flaws
B - Moderate	Moderate-quality randomized controlled trials Moderate-quality non-randomized studies
A - High	High-quality randomized controlled trials

GRADE

Certainty ratings	What it means
D - Very low	The true effect is probably markedly different from the estimated effect
C - Low	The true effect might be markedly different from the estimated effect
B - Moderate	The authors believe that the true effect is probably close to the estimated effect
A - High	The authors have a lot of confidence that the true effect is similar to the estimated effect

Implication of Strong and Weak Recommendations

	Strong (Level 1 = We Recommend)	Weak (Level 2 = We Suggest)
For patients	Many individuals would want course of action	Majority would want, but many would not
For clinicians	Many individuals should receive	Different choices appropriate for different patients Decision aids may be useful Shared decision consistent with values and preferences
For policy makers	Can be adapted as policy in most situations (or performance indicator)	Substantial debates Vary between regions

Recommendation and Suggestion

Blue colour: New item compared to 2015

Red colour: New grading compared to 2015

Main Topics to Cover

1. Sodium and fluid Restriction
2. Diuretics
3. Diabetes Management
4. Lipid Management
5. Heart failure Management
6. AF Management

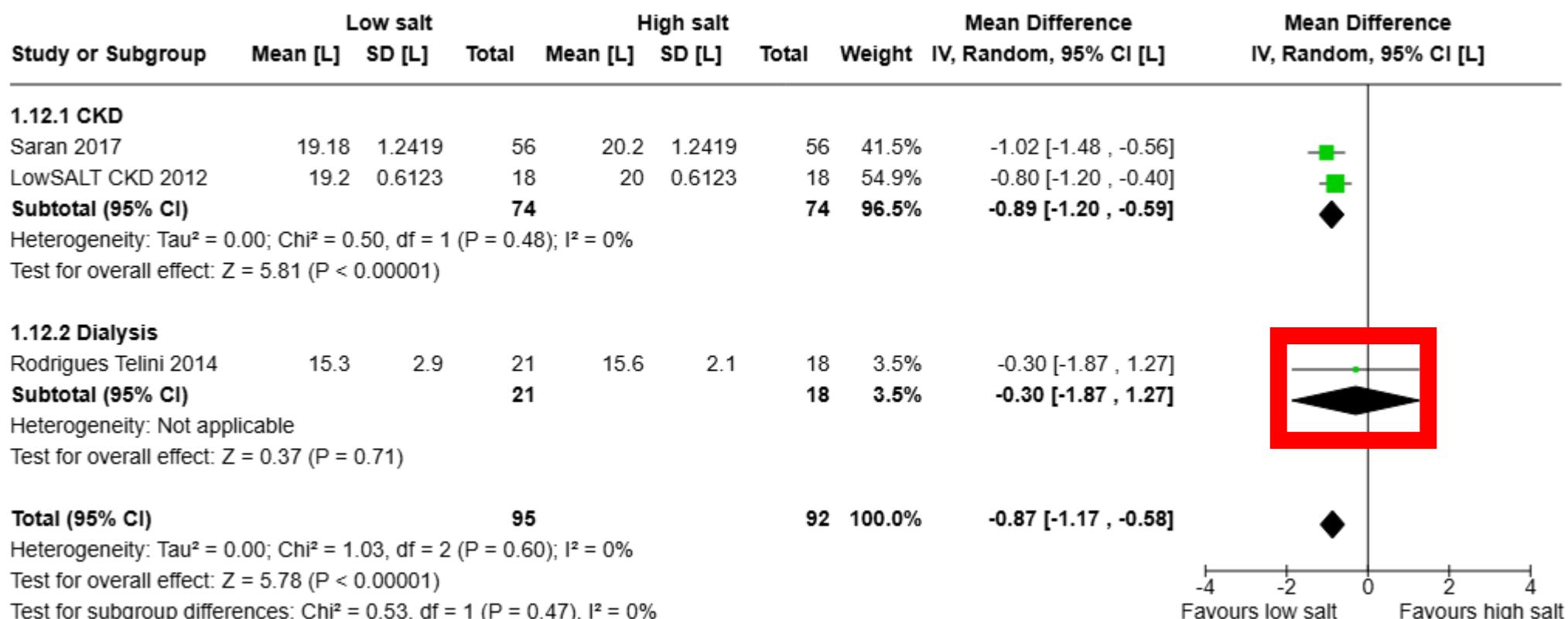
Sodium and Fluid Restriction

Cochrane Review

Evidence for sodium restriction to reduce extracellular fluid volume or edema is mostly available in the earlier stages of CKD (-0.87 L, 95% CI -1.17 to -0.58; 3 studies; 187 participants; low certainty evidence)

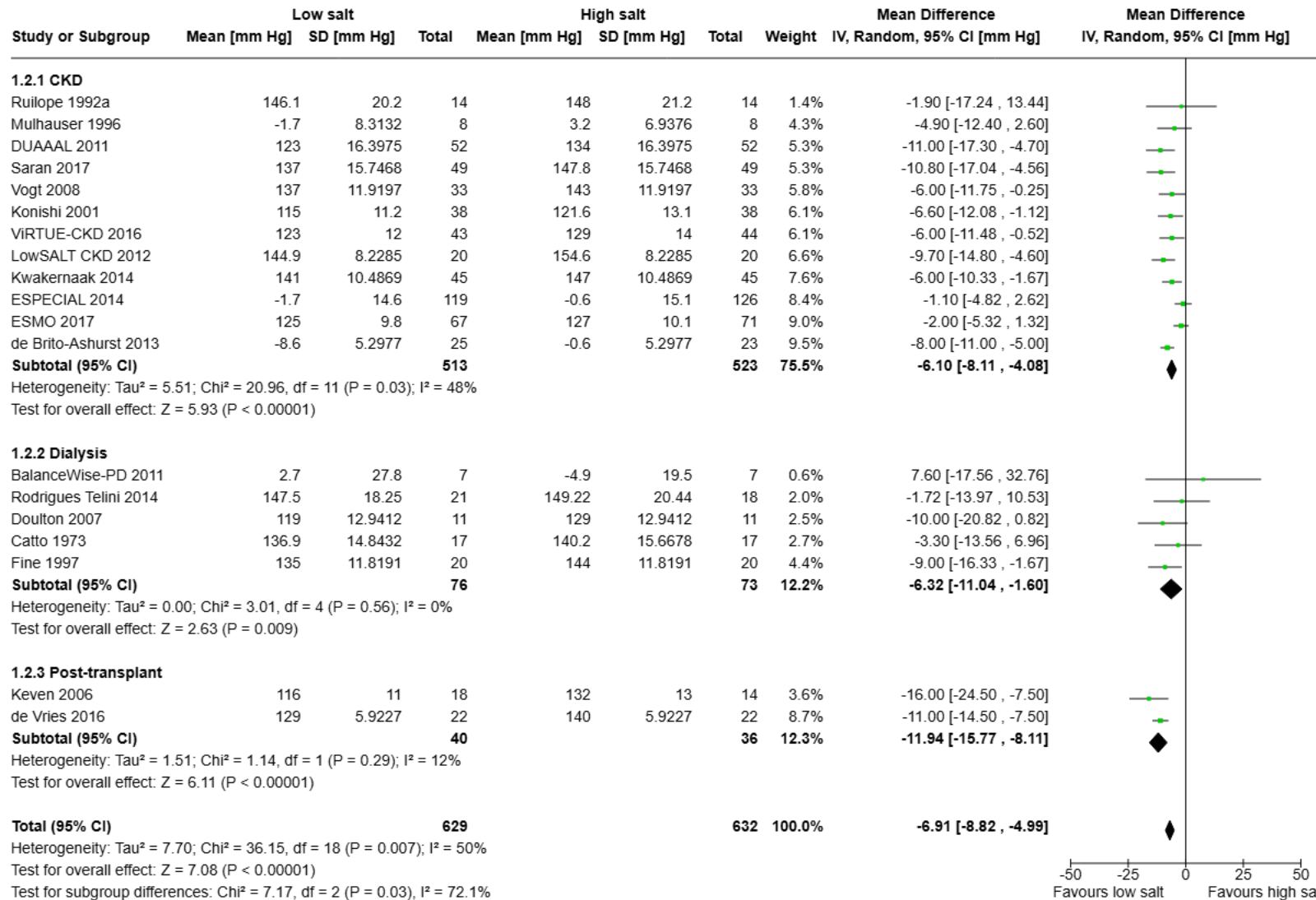
McMahon EJ, Campbell KL, Bauer JD, Mudge DW, Kelly JT. Altered dietary salt intake for people with chronic kidney disease. Cochrane Database Syst Rev 2021;6:CD010070

Extracellular Volume



McMahon EJ, Campbell KL, Bauer JD, Mudge DW, Kelly JT. Altered dietary salt intake for people with chronic kidney disease. Cochrane Database Syst Rev 2021;6:CD010070

Most Evidence on Systolic BP Only



ESC Consensus Statement



European Journal of Heart Failure (2024) 26, 730–741
doi:10.1002/ejhf.3244

POSITION PAPER

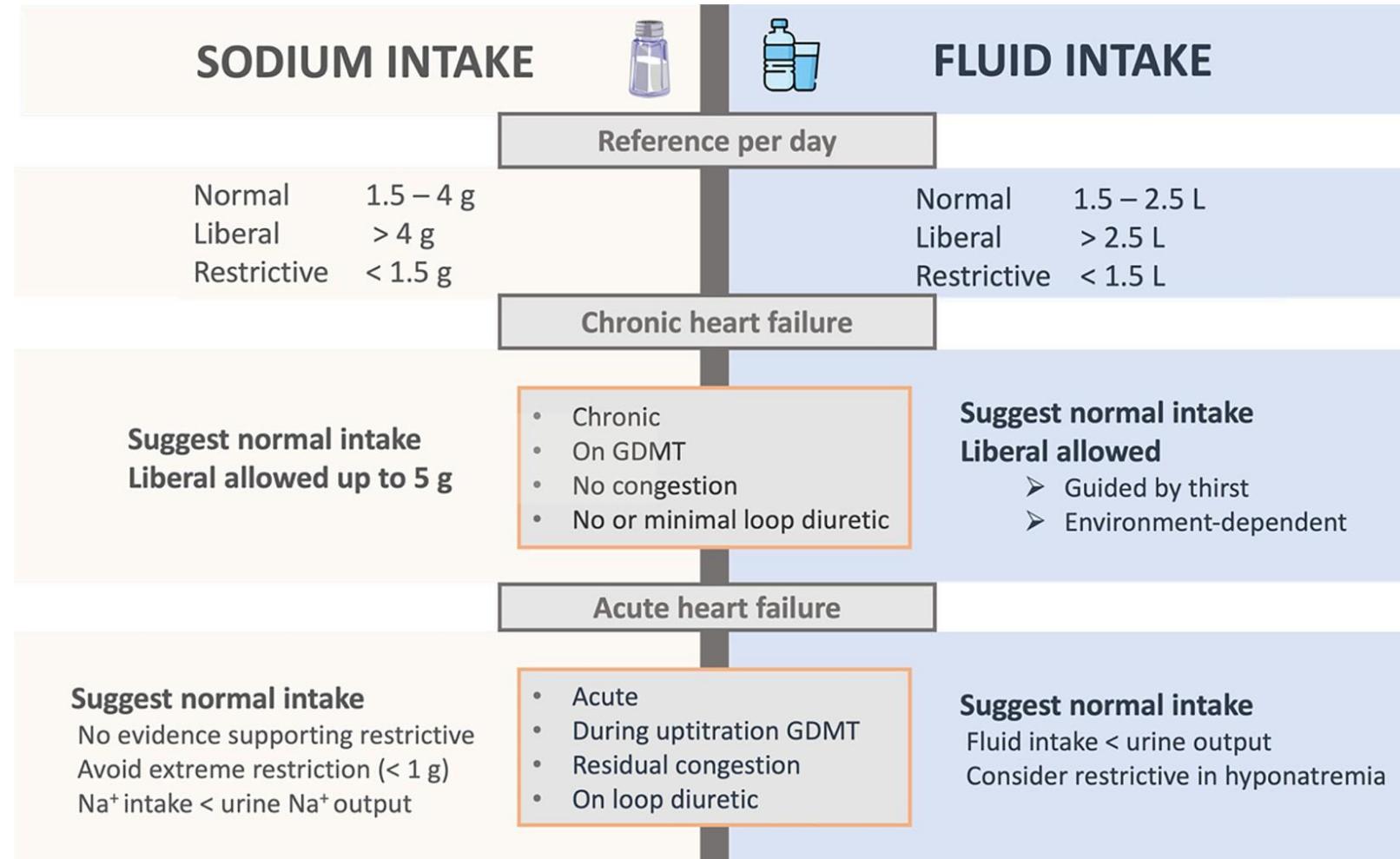
Dietary sodium and fluid intake in heart failure. A clinical consensus statement of the Heart Failure Association of the ESC

Wilfried Mullens^{1,2*}, Kevin Damman³, Sebastiaan Dhont^{1,2}, Debasish Banerjee⁴,
Antoni Bayes-Genis⁵, Antonio Cannata⁶, Ovidiu Chioncel⁷, Maja Cikes⁸,
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Mullens W, Damman K, Dhont S, Banerjee D, Bayes-Genis A, Cannata A, Chioncel O, Cikes M, Ezekowitz J, Flammer AJ, Martens P, Mebazaa A, Mentz RJ, Miró Ò, Moura B, Nunez J, Ter Maaten JM, Testani J, van Kimmenade R, Verbrugge FH, Metra M, Rosano GMC, Filippatos G. Dietary sodium and fluid intake in heart failure. A clinical consensus statement of the Heart Failure Association of the ESC. Eur J Heart Fail 2024;26:730-741

Study	Design	Patient population	Sample size	Intervention	Comparator	MACE
Fluid intake						
Holst <i>et al.</i>	Cross-over	HFrEF without clinical signs of congestion	74	Maximum fluid intake of 1.5 L/day	Fluid intake based on 30 ml/kg body weight/day	Readmission rate: NS
Albert <i>et al.</i>	Parallel-group, single-blind	ADHF, serum sodium ≤ 137 mg/dl	52	Maximum fluid intake of 1 L/day	Usual care discharge instructions and education	60-day mortality or readmission rate: NS
Sodium and fluid intake						
Aliti <i>et al.</i>	Parallel-group with blinded outcome assessments	Inpatients with ADHF, HFrEF	75	Sodium to 800 mg/day and fluid to 0.8 L/day during hospital stay	Unrestricted sodium and fluid intake	30-day readmission rate: NS
Machado d'Almeida <i>et al.</i>	Parallel-group with blinded outcome assessors	Inpatients with ADHF, HFpEF	53	Sodium to 800 mg/day and fluid to 0.8 L/day during hospital stay	Unrestricted sodium and fluid intake	30-day mortality or readmission rate: NS
Fabricio <i>et al.</i>	Single-blind	Patients hospitalized with ADHF	44	Low-sodium diet (3 g/day dietary salt) and fluid to 1 L/day	Normal-sodium diet (7 g/day salt) and fluid to 1 L/day	30-day readmission rate: NS
Sodium intake						
Hummel <i>et al.</i>	Single-blind, multicentre	Discharged from hospital with ADHF	66	Home-delivered sodium-restricted food (1500 mg/day sodium)	Usual care discharge instructions and education	30-day mortality or readmission rate: NS
Kalogeropoulos <i>et al.</i>	Double-blind	HFrEF with recent hospitalization on optimal GRMT	27	Sodium-restricted diet (1.5 g/day sodium)	Sodium-restricted diet (3 g/day sodium)	30-day readmission rate: NS
Ivey-Miranda <i>et al.</i>	Double-blind	HFrEF on optimal GRMT	70	Sodium-restricted diet (2 g/day sodium)	Sodium-restricted diet (3 g/day sodium)	30-day mortality or readmission rate: NS
Ezekowitz <i>et al.</i>	Multicentre open-label with blinded outcome assessments	Adult patients with chronic HF (NYHA class II-III) on optimal GRMT	806	Sodium-restricted diet (1.5 g/day sodium)	Usual care according to local guidelines	30-day mortality or readmission rate: NS



Mullens W, Damman K, Dhont S, Banerjee D, Bayes-Genis A, Cannata A, Chioncel O, Cikes M, Ezekowitz J, Flammer AJ, Martens P, Mebazaa A, Mentz RJ, Miró Ò, Moura B, Nunez J, Ter Maaten JM, Testani J, van Kimmenade R, Verbrugge FH, Metra M, Rosano GMC, Filippatos G. Dietary sodium and fluid intake in heart failure. A clinical consensus statement of the Heart Failure Association of the ESC. Eur J Heart Fail 2024;26:730-741

Main Messages from ESC

- Limitation of salt intake to no more than 5 g/day in patients with heart failure
- Contemplating fluid restriction of 1.5–2 L/day only in selected patients

Mullens W, Damman K, Dhont S, Banerjee D, Bayes-Genis A, Cannata A, Chioncel O, Cikes M, Ezekowitz J, Flammer AJ, Martens P, Mebazaa A, Mentz RJ, Miró Ò, Moura B, Nunez J, Ter Maaten JM, Testani J, van Kimmenade R, Verbrugge FH, Metra M, Rosano GMC, Filippatos G. Dietary sodium and fluid intake in heart failure. A clinical consensus statement of the Heart Failure Association of the ESC. Eur J Heart Fail 2024;26:730-741

More Recent RCT in Ambulatory Heart Failure

504 patients with chronic HF for at least 6 months

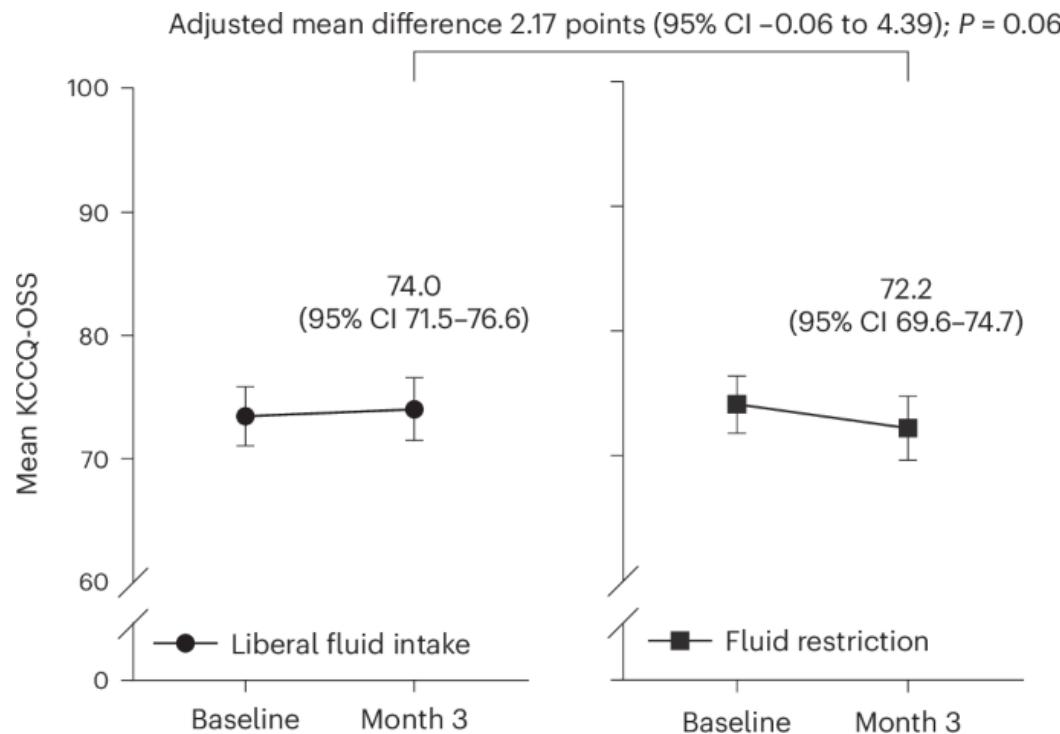
Randomized to receive lifestyle advice recommending either

- fluid restriction (<1.5 L/day), or
- liberal fluid intake (no maximum)

Herrmann JJ, Brunner-La Rocca HP, Baltussen LEHJM, Beckers-Wesche F, Bekkers SCAM, Bellersen L, van Eck JWM, Hassing HC, Jaarsma T, Linssen GCM, Pisters R, Sanders-van Wijk S, Verdijk MHI, Handoko ML, van der Meer P, Verbrugge FH, Januzzi JL Jr, Bayés-Genís A, Nieuwlaat R, Rodwell L, Gommans DHF, van Kimmenade RRJ. Liberal fluid intake versus fluid restriction in chronic heart failure: a randomized clinical trial. *Nat Med* 2025;31:2062-2068

Primary Outcome

Kansas City Cardiomyopathy Questionnaire summary score, 72.2 vs. 74.0 points (no difference)



Fluid Restriction in Heart Failure

ACC/AHA guideline recommendation

- Consider fluid restriction only for those with advanced heart failure and hyponatremia (Grade 2B)



SALT, FLUID MONITORING AND MANAGEMENT

- We suggest restricting dietary salt < 5 g/day (sodium < 2 g/day) and fluid intake to manage volume overload in PD patients (**2C**; downgraded from 1C).

Diuretics

Landmark Study: frusemide in patients on PD

61 patients newly started on **CAPD**

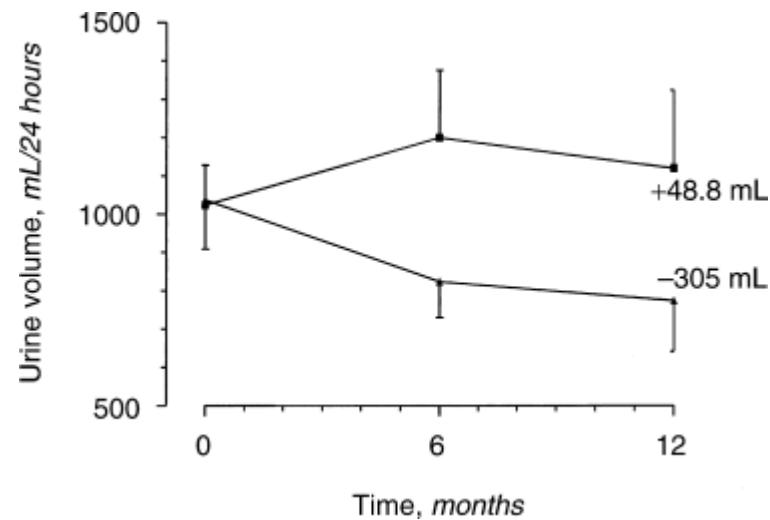
Eligibility: older than 16, and 10 days after CAPD training

Exclusion: had received more than two weeks of hemodialysis and those with a failing kidney transplant

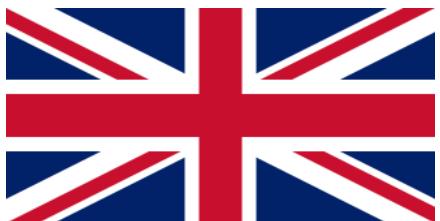
Randomized in a 1:1 ratio to open-label frusemide 250 mg daily or no diuretics

Medcalf JF, Harris KP, Walls J. Role of diuretics in the preservation of residual renal function in patients on continuous ambulatory peritoneal dialysis. *Kidney Int* 2001;59:1128-33

Frusemide and CAPD: 12 months follow up



- Increase in urine volume
- Increase in urinary sodium excretion
- Less increase in total body weight
- No effect on residual kidney function



Medcalf JF, Harris KP, Walls J. Role of diuretics in the preservation of residual renal function in patients on continuous ambulatory peritoneal dialysis. *Kidney Int* 2001;59:1128-33

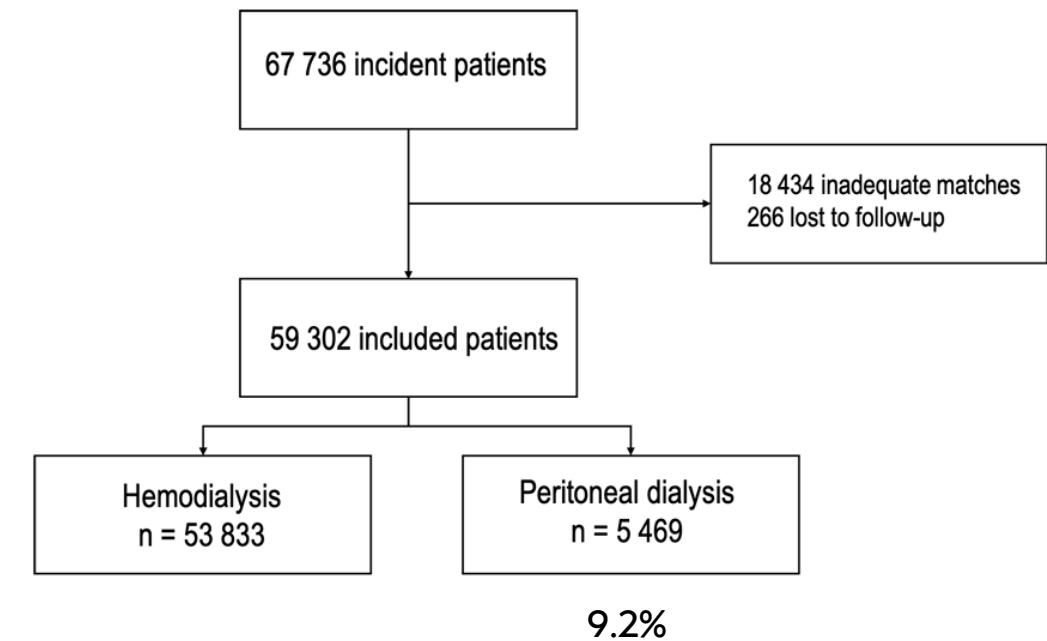
REIN registry Study: diuretics in patients on PD

Observational study of adult patients (> 18 years) who started dialysis between 2009 and 2015

Classified by loop diuretics

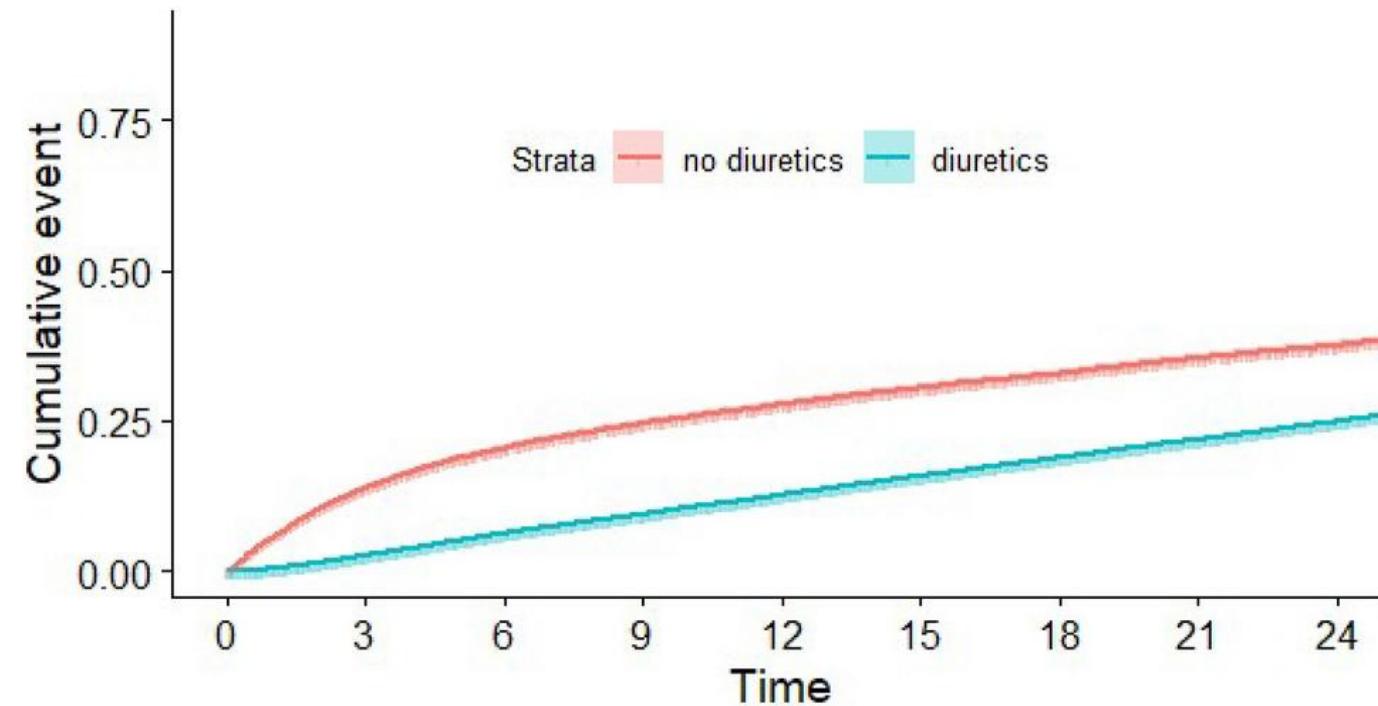
exposure: < 5%, 5–50%, 50–80%

or > 80% over the observation period



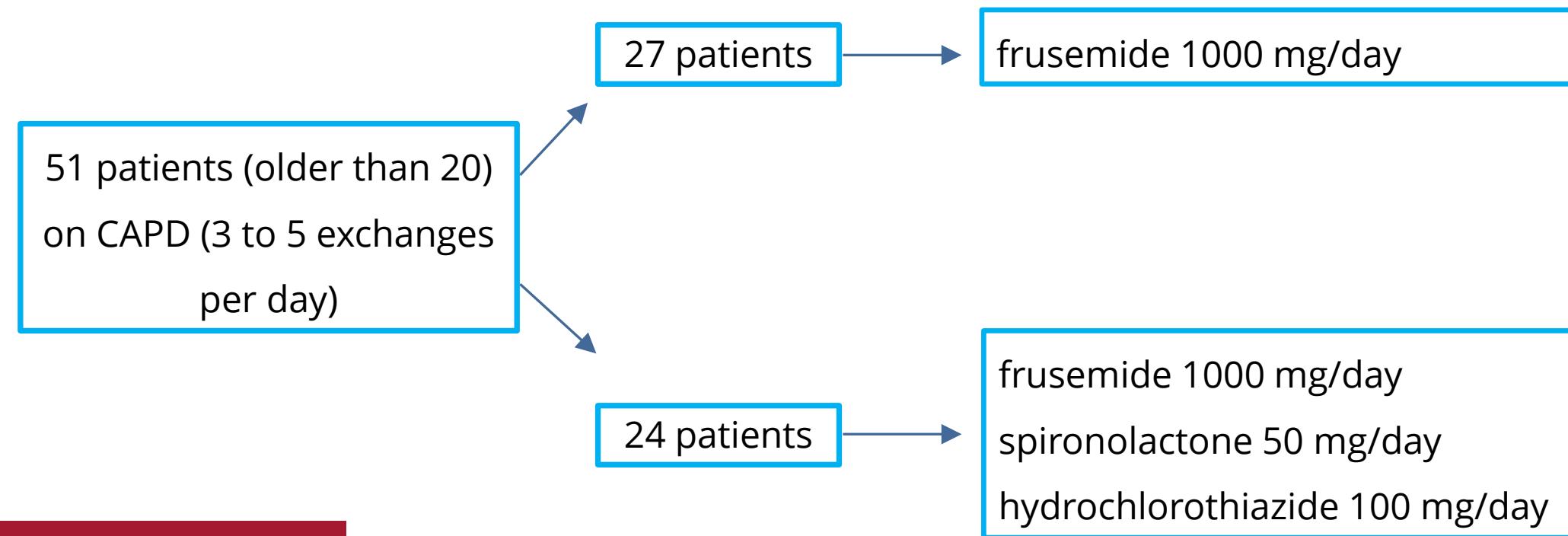
Ingwiller M, Bozman DF, Florens N, Cerasuolo D, Vigneau C, Couchoud C, Hannedouche T. Diuretics and mortality reduction in incident dialysis patients: a two-year observational study. *Sci Rep* 2024;14:27447

Significantly lower 2-year mortality (propensity score matching)



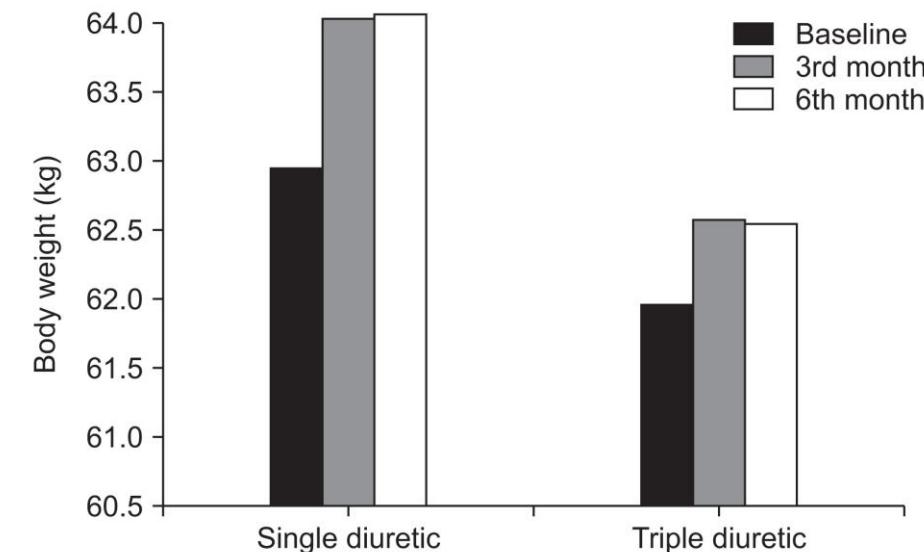
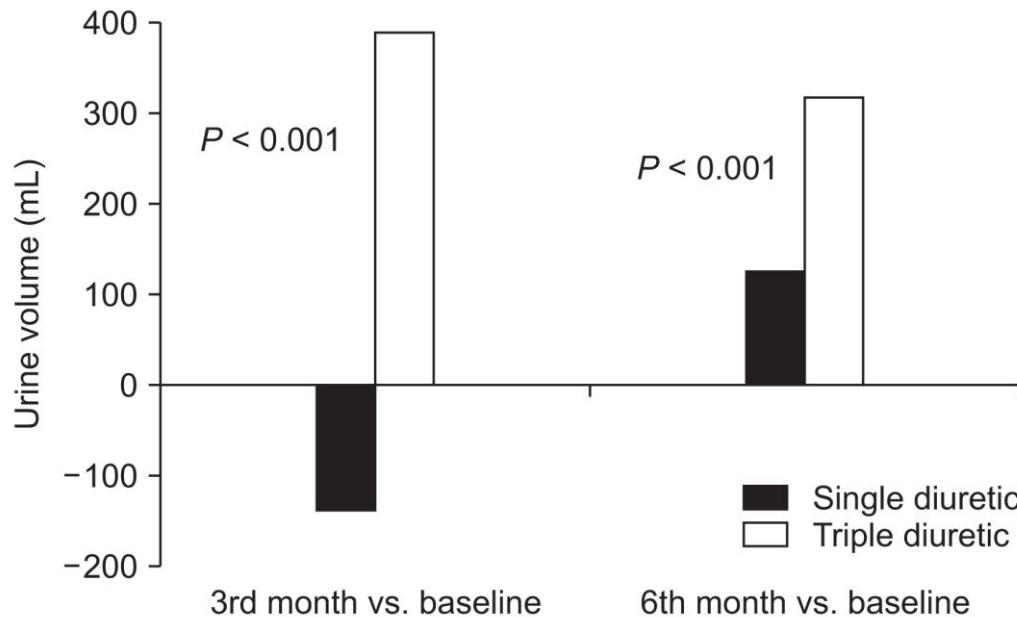
Ingwiller M, Bozman DF, Florens N, Cerasuolo D, Vigneau C, Couchoud C, Hannedouche T. Diuretics and mortality reduction in incident dialysis patients: a two-year observational study. *Sci Rep* 2024;14:27447

RCT: triple diuretics in patients on PD



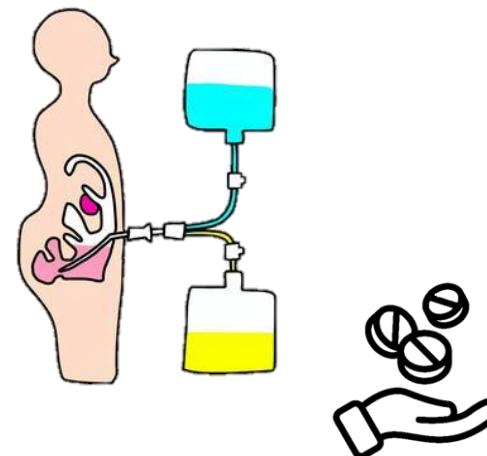
Witoon R, Yongsiri S, Buranaburidej P, Nanna P. Efficacy of triple diuretic treatment in continuous ambulatory peritoneal dialysis patients: a randomized controlled trial. *Kidney Res Clin Pract* 2019;38:108-115

Better urine output and volume control



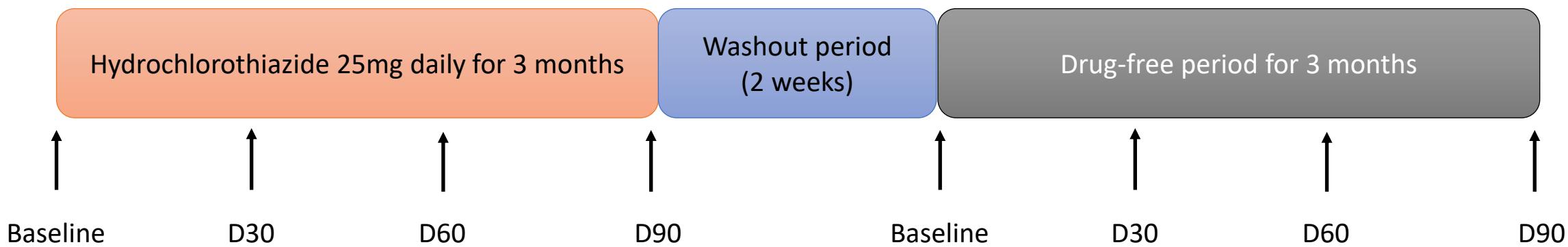
Witoon R, Yongsiri S, Buranaburidej P, Nanna P. Efficacy of triple diuretic treatment in continuous ambulatory peritoneal dialysis patients: a randomized controlled trial. *Kidney Res Clin Pract* 2019;38:108-115

Single Centre Study for Thiazide



Cross-over study

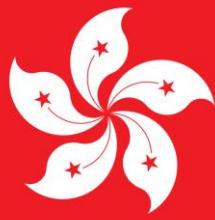
- 10 adult prevalent patients on PD with uncontrolled hypertension (SBP 140 or DBP >90) with residual kidney function (> 500ml/day)

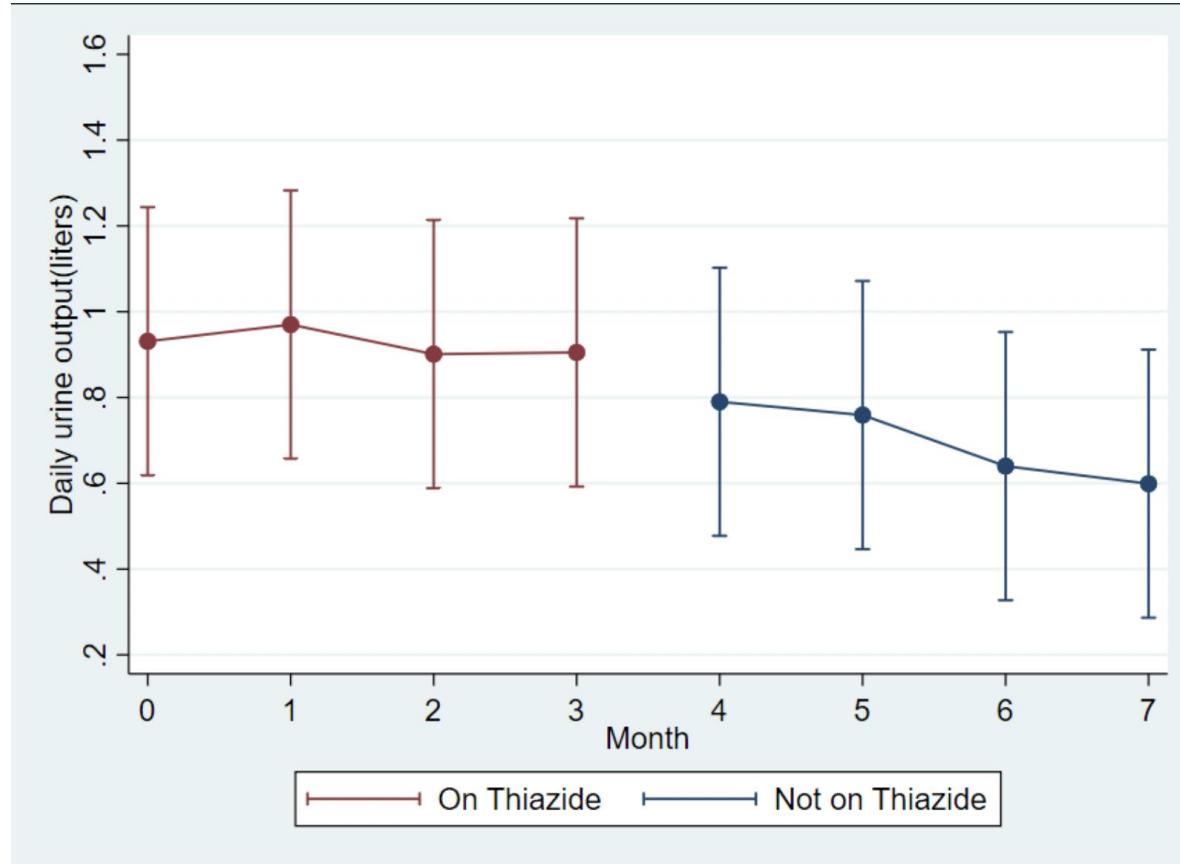


Assessments: BP, BW, BH, BCM (by bioimpedance spectroscopy), electrolytes, 24-hour urine volume, CrCl, Na

Unpublished Results

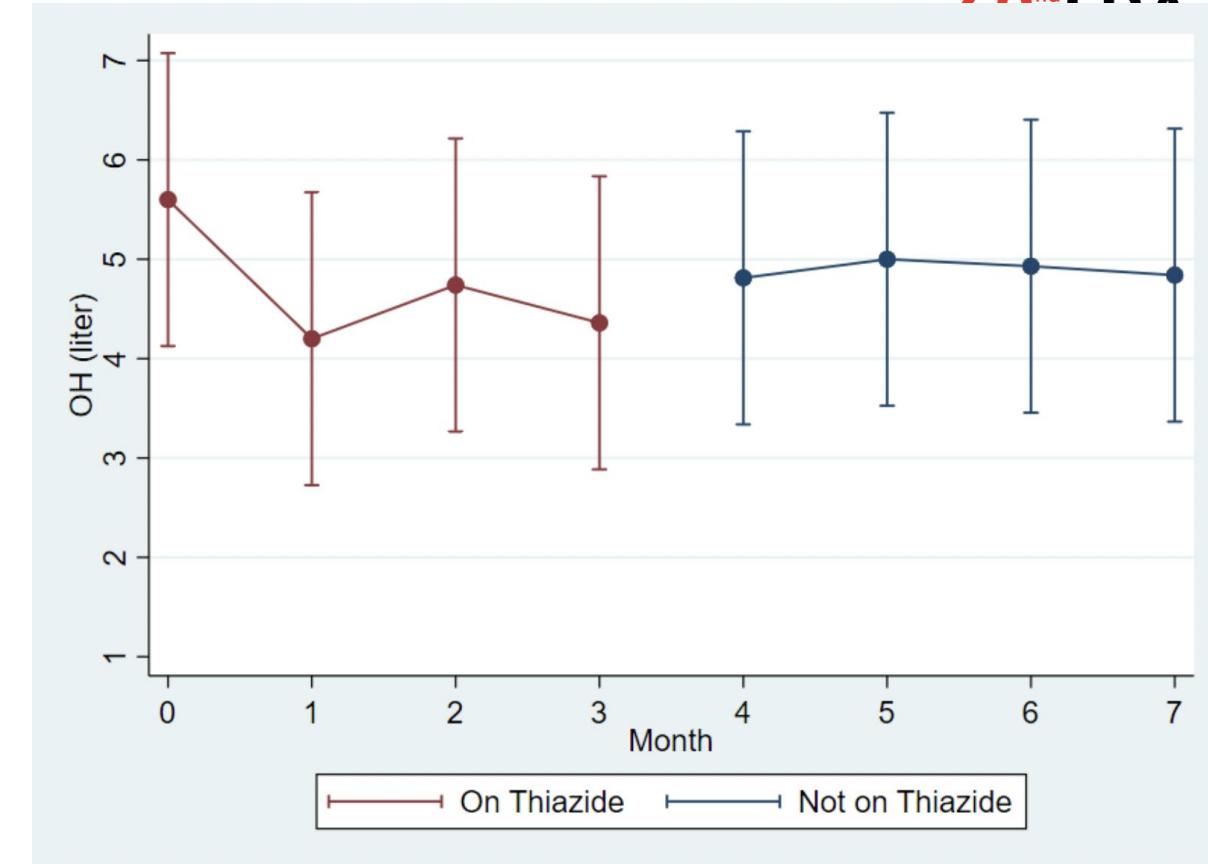
	$n = 10$
Age	64.4 ± 11.41
Male gender	7 (70%)
Primary diagnosis	
- Diabetes mellitus	3 (30%)
- Hypertension	2 (20%)
- Glomerulonephritis	3 (30%)
- Unknown	2 (20%)
Loop diuretics use	10 (100%)
Systolic blood pressure (mmHg)	164 ± 9
Diastolic blood pressure (mmHg)	87 ± 14
Daily urine volume (L)	1.04 ± 0.42





Decline in urine output

On Thiazide	-0.03 L /3 months (-0.1 L/year)
Not on Thiazide	-0.31 L /3 months (-1.24 L/year)



Overhydration Difference

On Thiazide	↓ 1.24L
Not on Thiazide	↑ 0.48L

Key (Unpublished) Results of Thiazide in patients receiving PD

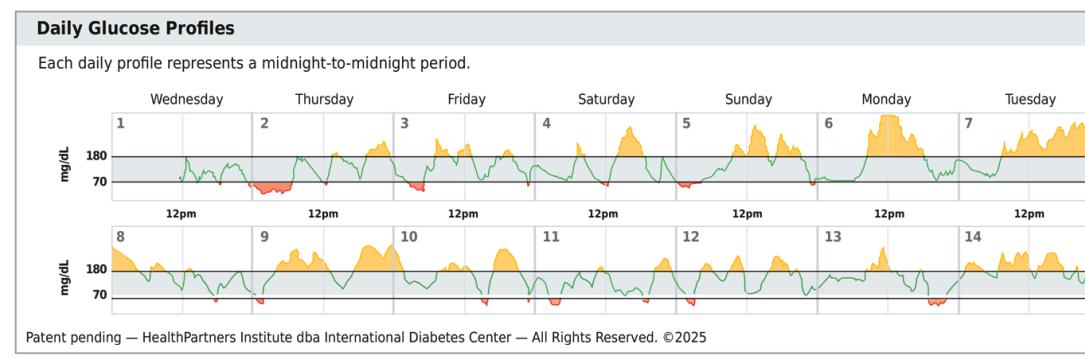
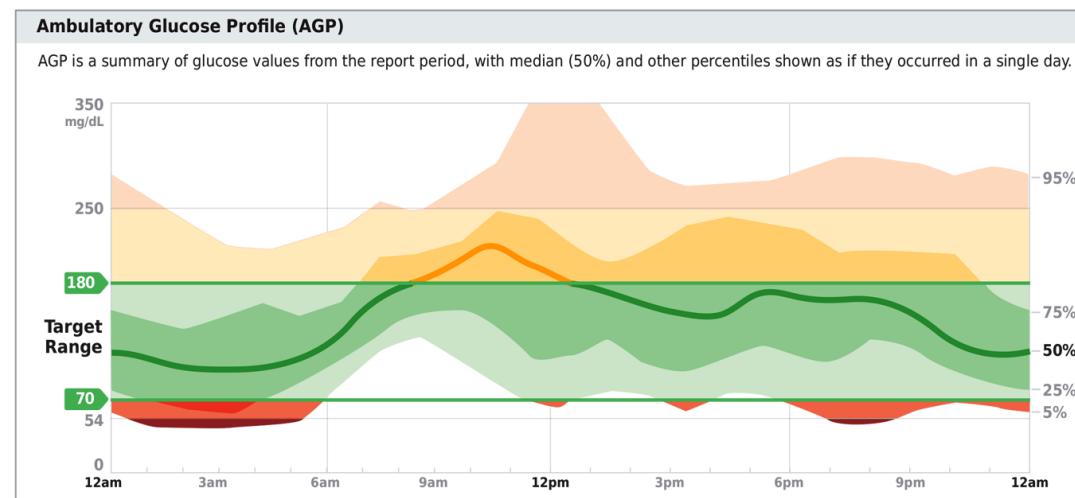
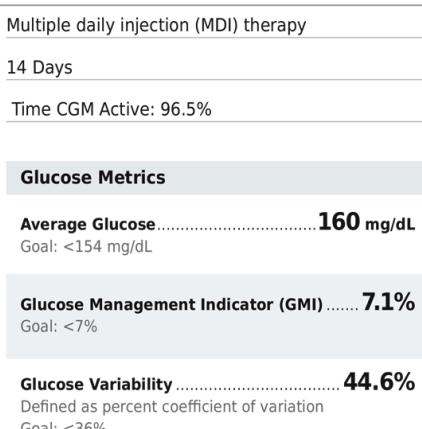
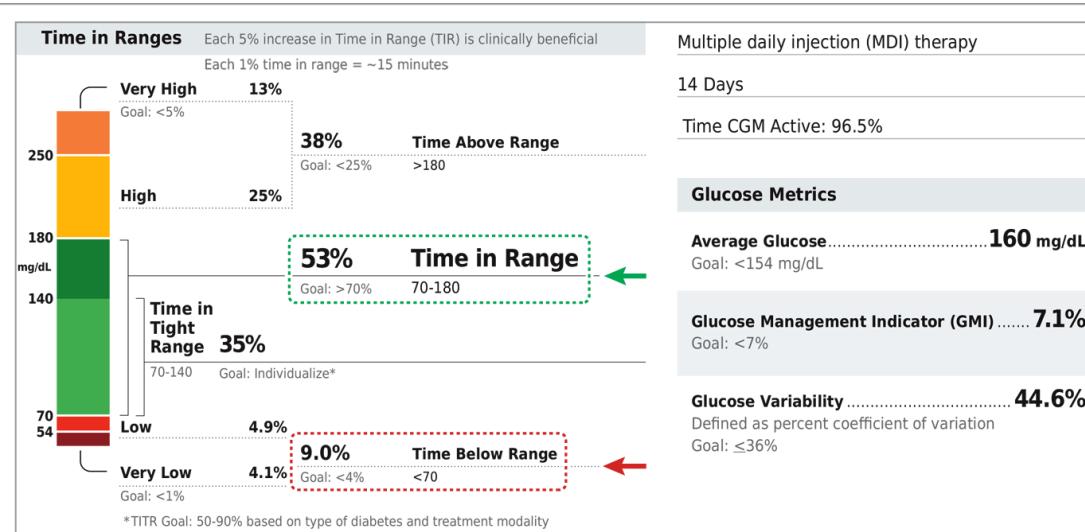
	On Thiazide	Not on Thiazide	Difference
Blood pressure	SBP: ↓ 13.0 ± 10.4	↑ 10.4 ± 14.0	23.4 mmHg
	DBP: ↓ 13.0 ± 10.4	↑ 2.3 ± 13.1	15.2 mmHg
Decline in urine output	-0.03 L /3 months (-0.10 L /year)	-0.31 L /3 months (-1.24 L /year)	-0.28 L /3 months (-1.14 L /year)
Overhydration	↓ 1.24L	↑ 0.48L	1.72L



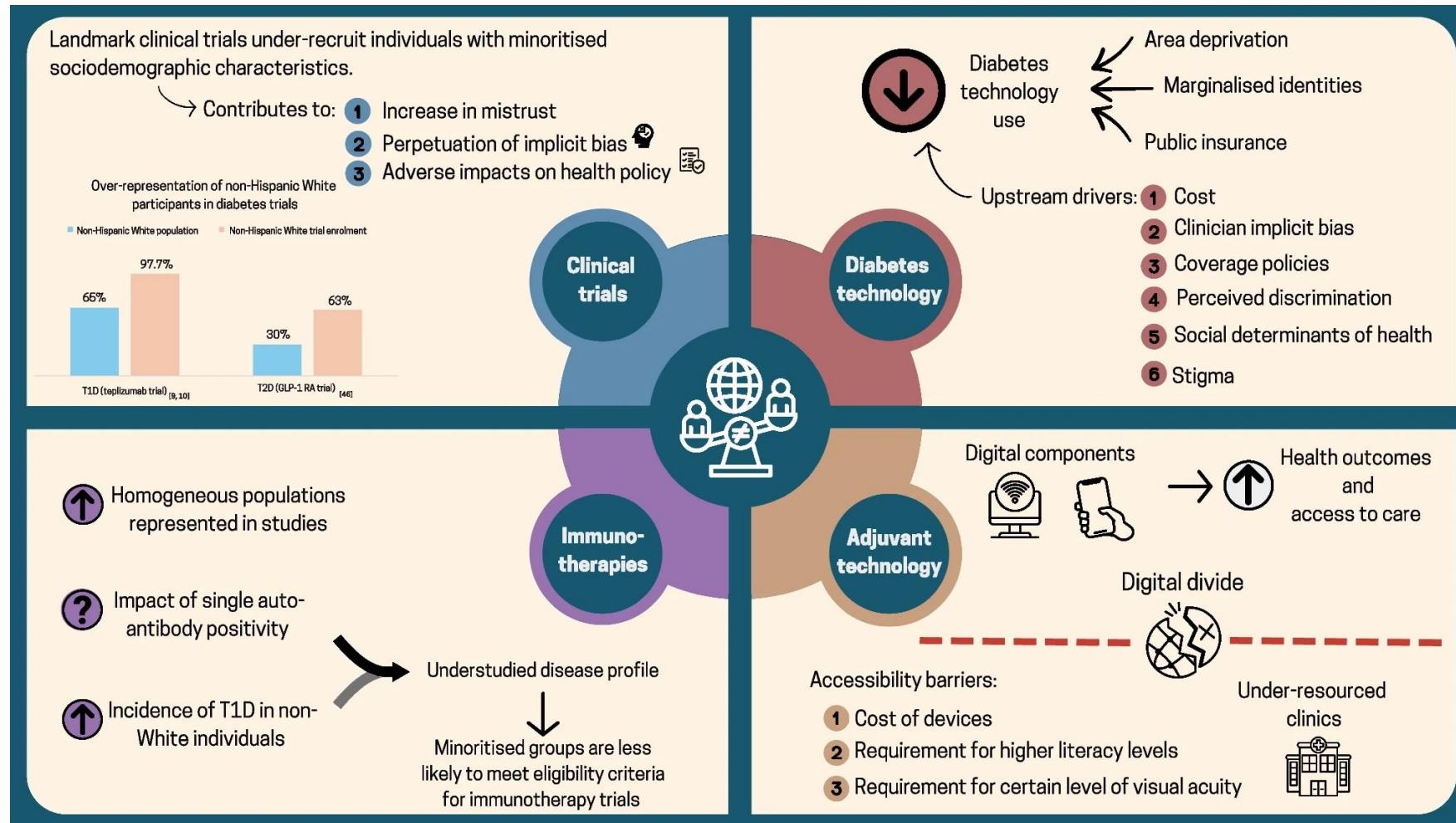
SALT, FLUID MONITORING AND MANAGEMENT

- We recommend maximizing loop diuretics to manage volume overload in patients on PD with RKF or daily urine volume more than 100 mL (1B).
- We suggest adding thiazide or thiazide-like diuretics as an adjunctive therapy to manage fluid overload (2B).

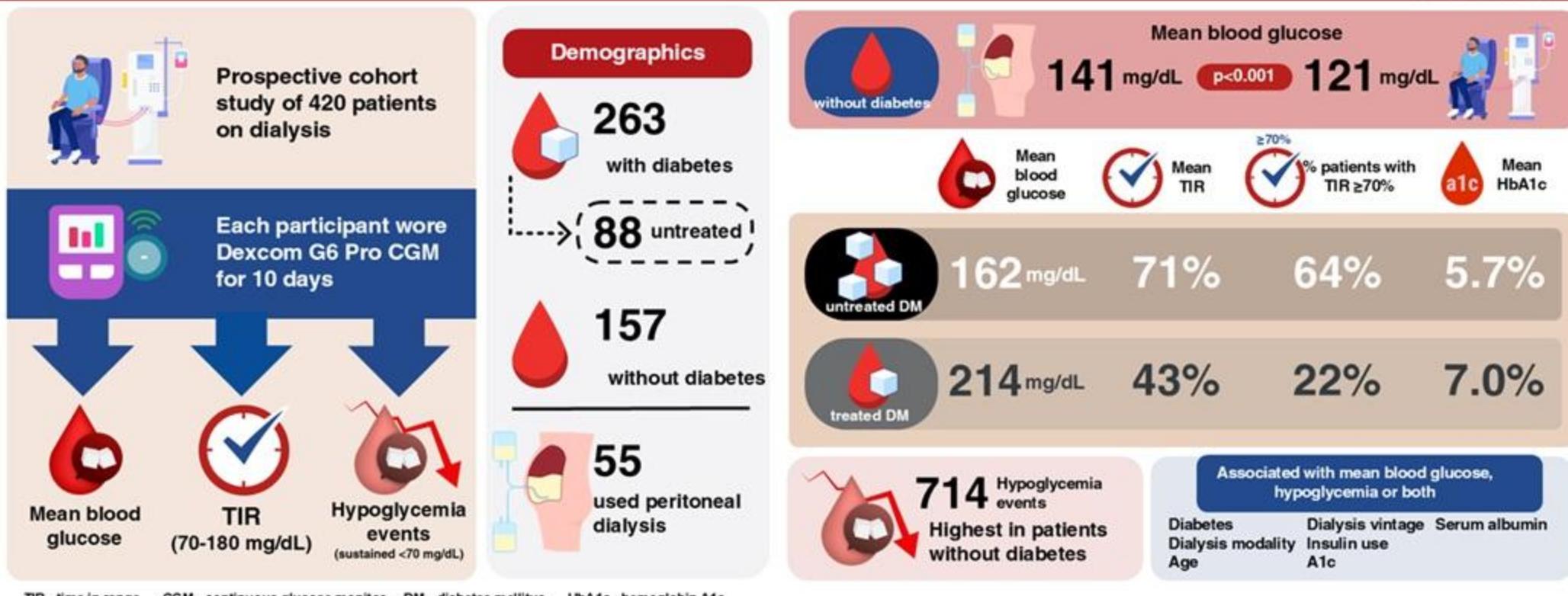
Continuous Glucose Monitoring



Bergenstal RM, Martens TW, Beck RW. Continuous glucose monitoring. JAMA 2025 Epub ahead of print



Glycemia Assessed by Continuous Glucose Monitoring among People Treated with Maintenance Dialysis



Conclusions: In maintenance dialysis, CGM frequently identified both hyperglycemia and hypoglycemia that may not be clinically evident. In particular, hyperglycemia was common with peritoneal dialysis, patients with untreated diabetes maintain a diabetic glycemic profile, and patients with treated diabetes rarely meet contemporary CGM-based treatment targets.

Ian H. de Boer, Lisa D. Anderson, Nathaniel K. Ashford, et al. *Glycemia Assessed by Continuous Glucose Monitoring among People Treated with Maintenance Dialysis*. JASN doi: 10.1681/ASN.0000000693. Visual Abstract by Carlo Nemesio B. Trinidad, MD

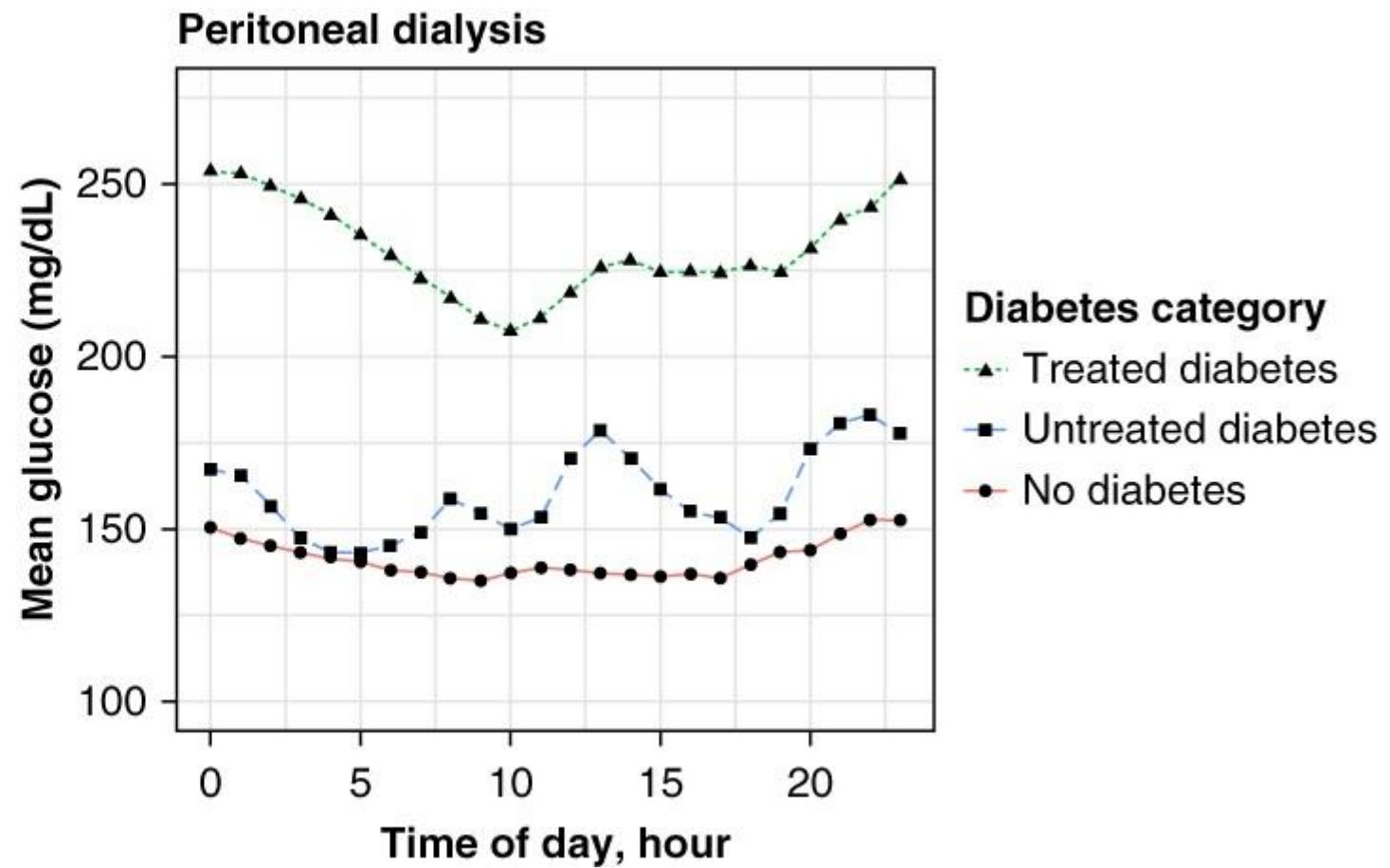
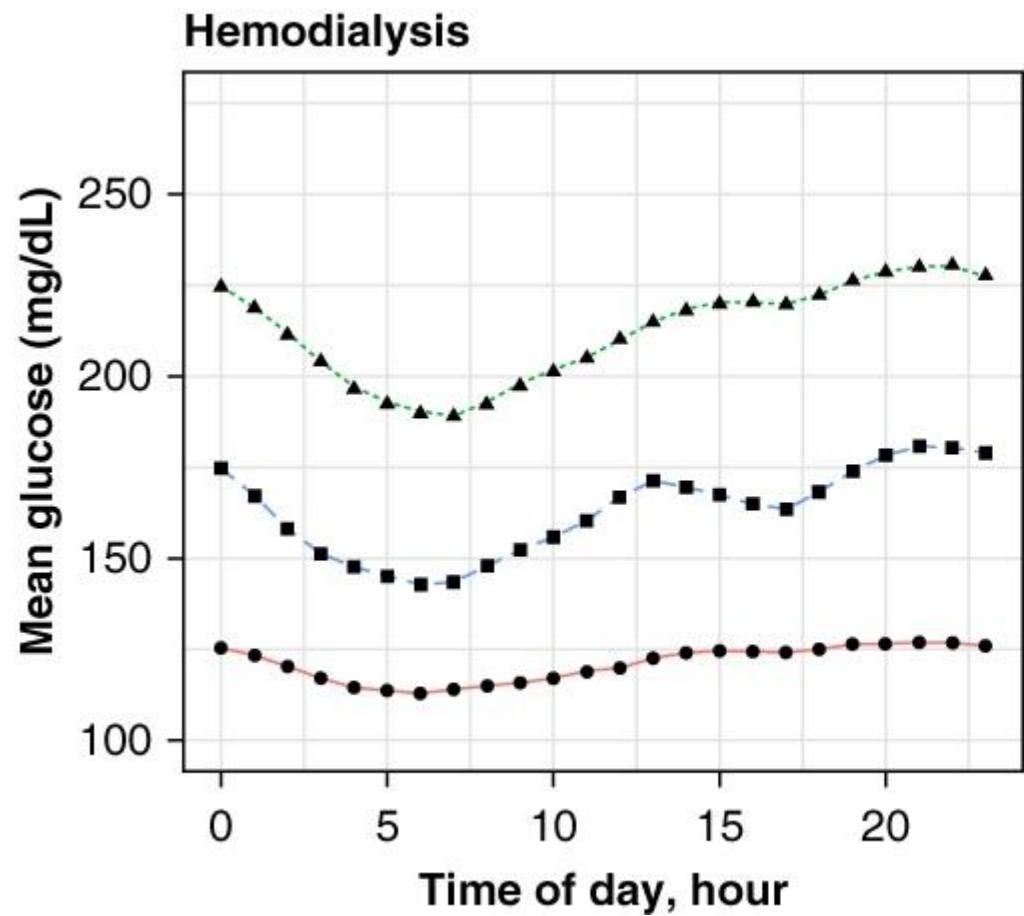
BLOSSOM Study

- 420 individuals (263 with diabetes and 157 without diabetes) treated with dialysis (hemodialysis, $n=365$; peritoneal dialysis, $n=55$)
- CGM identified episodes of hyperglycemia and hypoglycemia that were not otherwise clinically evident, even when HbA1c $< 7\%$ (53 mmol/mol)



de Boer IH, Anderson LD, Ashford NK, Ayers E, Bansal N, Hall YN, Hirsch IB, Hoofnagle AN, Hsu S, Jones E, Lidgard B, Limonte CP, Linke LJ, Marnell CC, Mayeda L, McNamara E, Mehrotra R, Pesenson A, Porter JM, Rivara MB, Roberts GV, Shanaman B, Trikudanathan S, Watnick S, Wilkens KG, Zelnick LR. Glycemia assessed by continuous glucose monitoring among people treated with maintenance dialysis. *J Am Soc Nephrol* 2025;36:1798-1810

Diurnal Pattern





GLUCOSE MANAGEMENT

- We suggest using continuous glucose monitoring (CGM) as an adjunct to HbA1c in patients on PD with diabetes to achieve individualized glycemic goal (2C).
- For PD patients who use CGM for glucose monitoring, we consider an individualized target of time in range (3.9–10 mmol/L) (70–180 mg/dL) over 50% appropriate (Not Graded).

Lipid



LIPID MANAGEMENT – SECONDARY PREVENTION

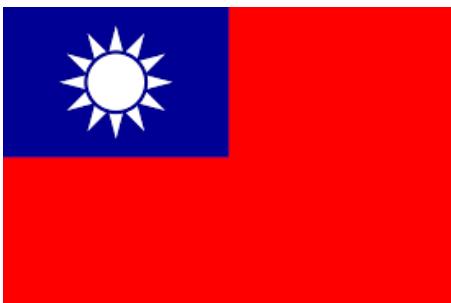
Secondary prevention of ASCVD

- Supported statin use in observational studies
- Most retrospective studies are propensity-matched
- Meta-analysis published in 2023

Statin after PCI for acute myocardial infarction

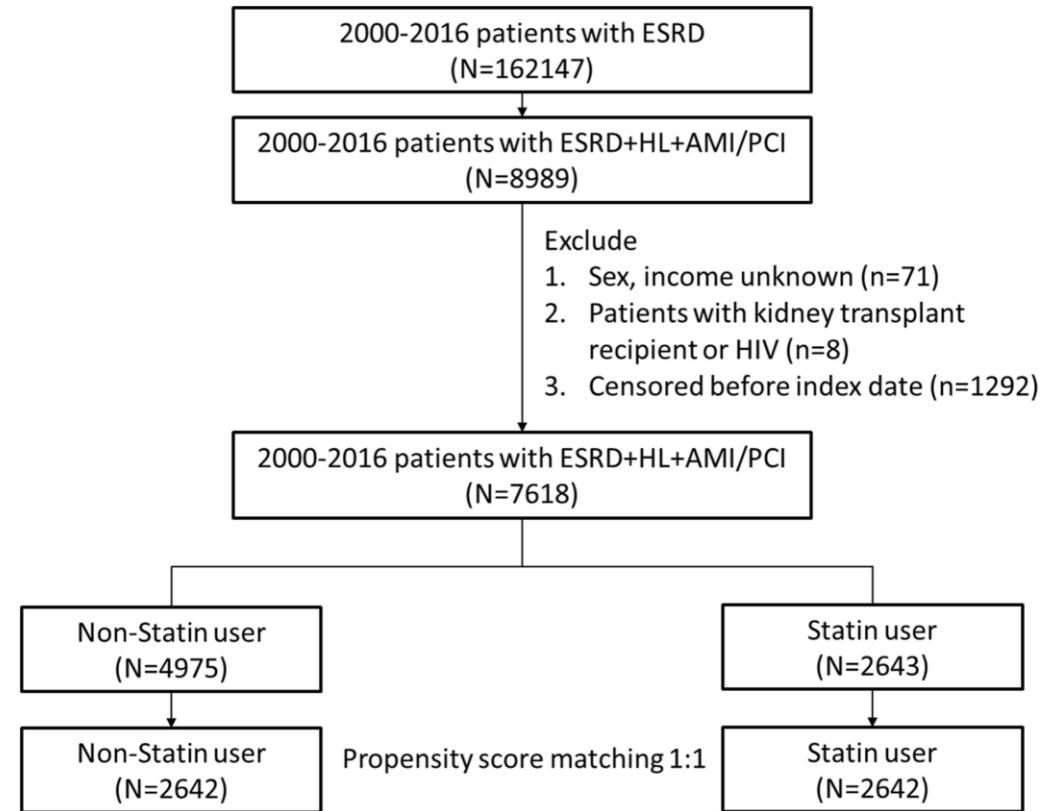
National Health Insurance of Taiwan in 2000–2016

- Patients on dialysis and with hyperlipidemia, with AMI and receiving PCI
- Propensity score matching



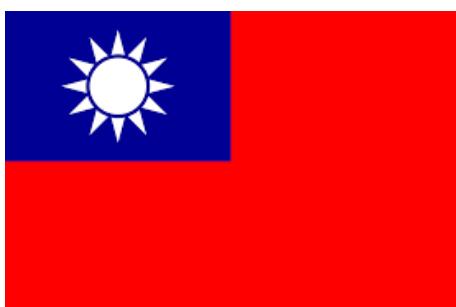
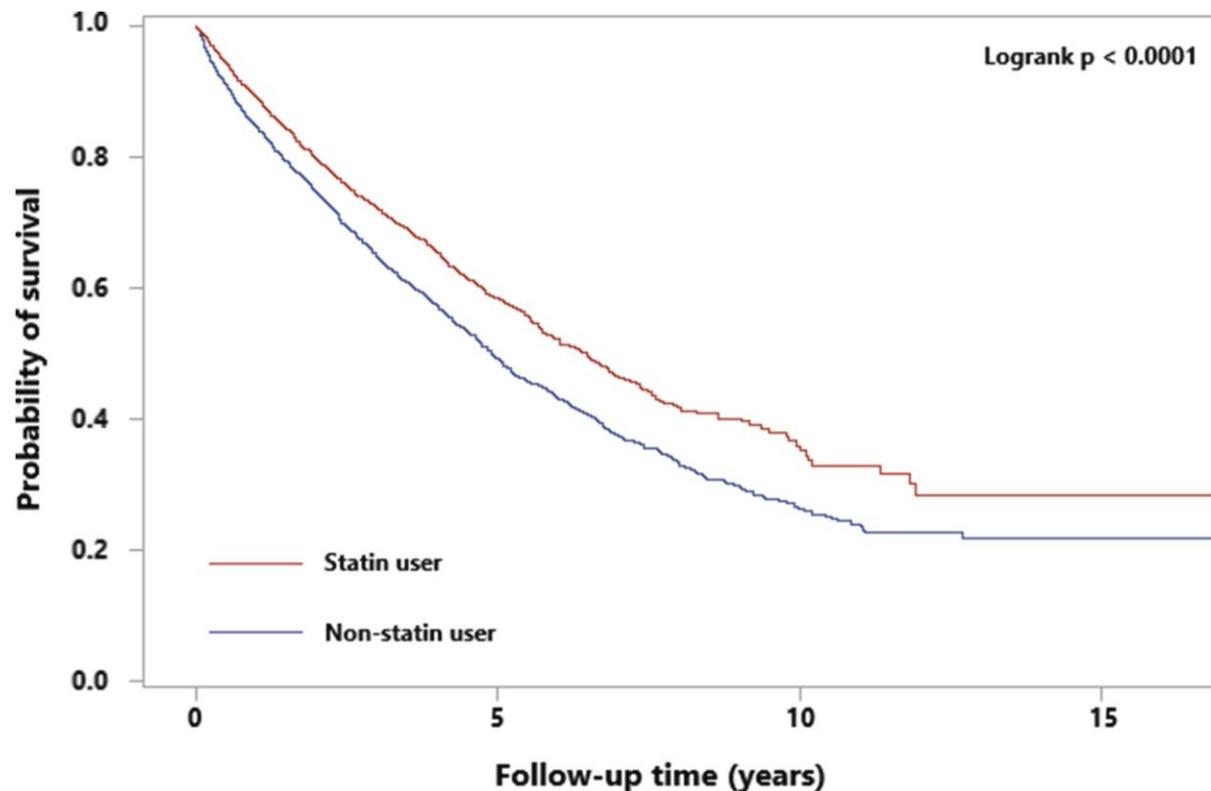
Yeh YT, Sung FC, Tsai CF, Hsu CC, Tsai WC, Hsu YH. Statin therapy associated mortality in hyperlipidemic dialysis patients with percutaneous coronary intervention for acute myocardial infarction, a retrospective cohort study. *Heliyon* 2024;10:e39906

Statin after PCI for acute myocardial infarction



Yeh YT, Sung FC, Tsai CF, Hsu CC, Tsai WC, Hsu YH. Statin therapy associated mortality in hyperlipidemic dialysis patients with percutaneous coronary intervention for acute myocardial infarction, a retrospective cohort study. *Heliyon* 2024;10:e39906

Statin: All-cause mortality reduced by 23%



Yeh YT, Sung FC, Tsai CF, Hsu CC, Tsai WC, Hsu YH. Statin therapy associated mortality in hyperlipidemic dialysis patients with percutaneous coronary intervention for acute myocardial infarction, a retrospective cohort study. *Helion* 2024;10:e39906

Statin after atherosclerotic cardiovascular disease

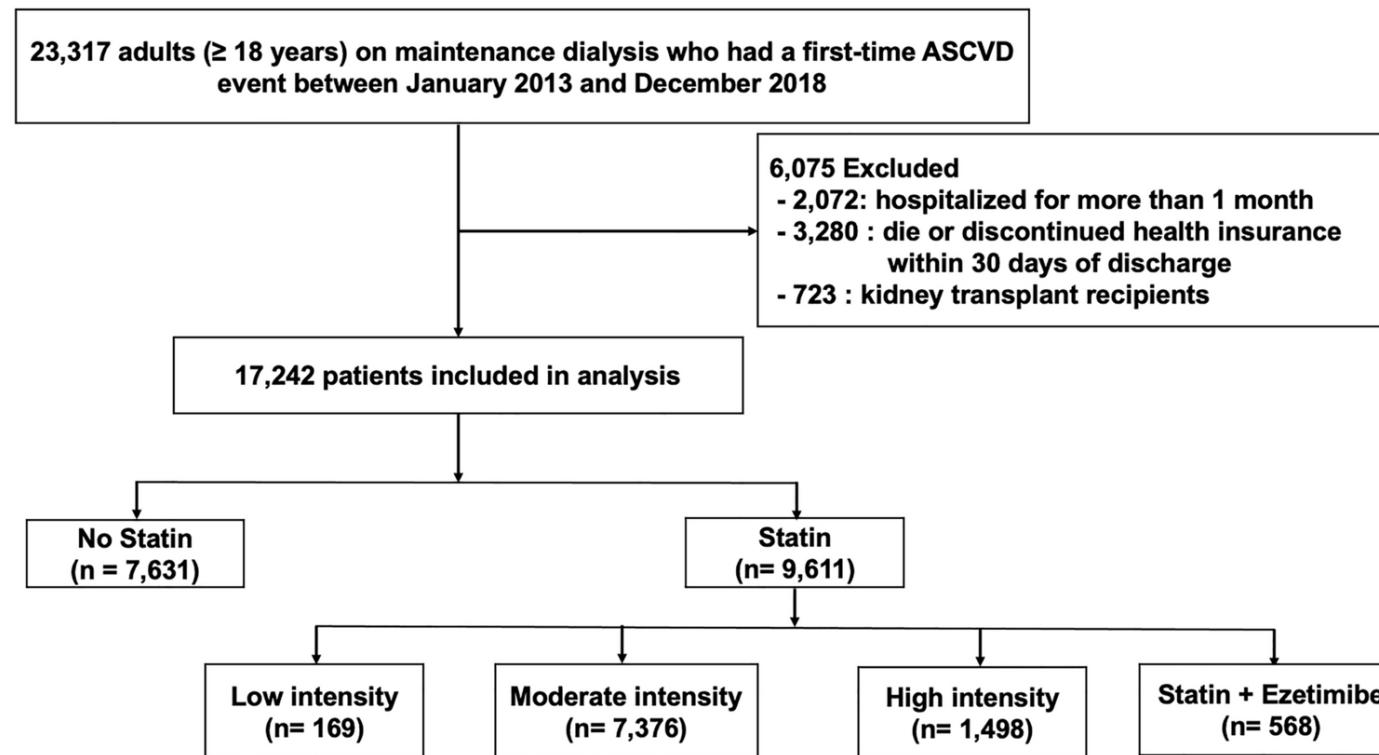
Korean NHIS (Nationwide Health Insurance Service) database

- 17,242 patients on dialysis who experienced a first-time ASCVD event
- 9611 patients (55.7%) received statin



Lee M, Choi WJ, Lee Y, Lee K, Park MW, Myong JP, Kim DW. Association between statin therapy and mortality in patients on dialysis after atherosclerotic cardiovascular diseases. *Sci Rep* 2023;13:10940

Statin after atherosclerotic cardiovascular disease



Lee M, Choi WJ, Lee Y, Lee K, Park MW, Myong JP, Kim DW. Association between statin therapy and mortality in patients on dialysis after atherosclerotic cardiovascular diseases. Sci Rep 2023;13:10940

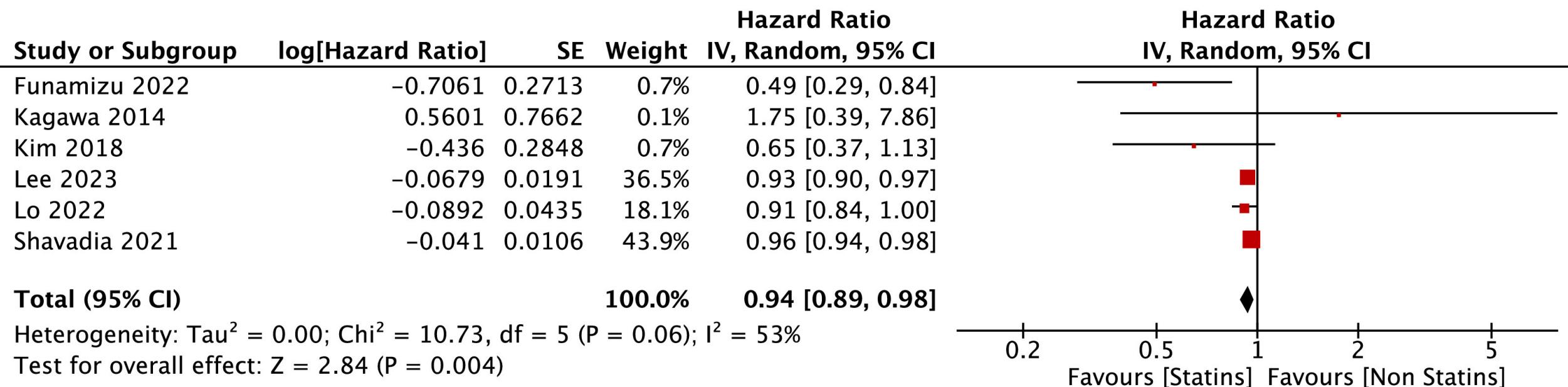
Statin reduced all-cause mortality

	Unadjusted		Adjusted for age and sex		Fully adjusted	
	HR (95% CI)	p value	HR (95% CI)	p value	HR (95% CI)	p value
All-cause mortality						
No statin	1 (Reference)		1 (Reference)		1 (Reference)	
Statin						
Overall	0.98 (0.94–1.03)	0.454	0.96 (0.92–1.01)	0.086	0.92 (0.88–0.97)	0.0009
CHD	1.07 (1.01–1.14)	0.035	1.01 (0.95–1.08)	0.683	0.97 (0.92–1.04)	0.411
CVA	0.87 (0.78–0.96)	0.006	0.89 (0.81–0.99)	0.034	0.88 (0.80–0.98)	0.021
PAD	1.09 (0.98–1.22)	0.107	1.10 (0.98–1.22)	0.099	1.03 (0.92–1.15)	0.598



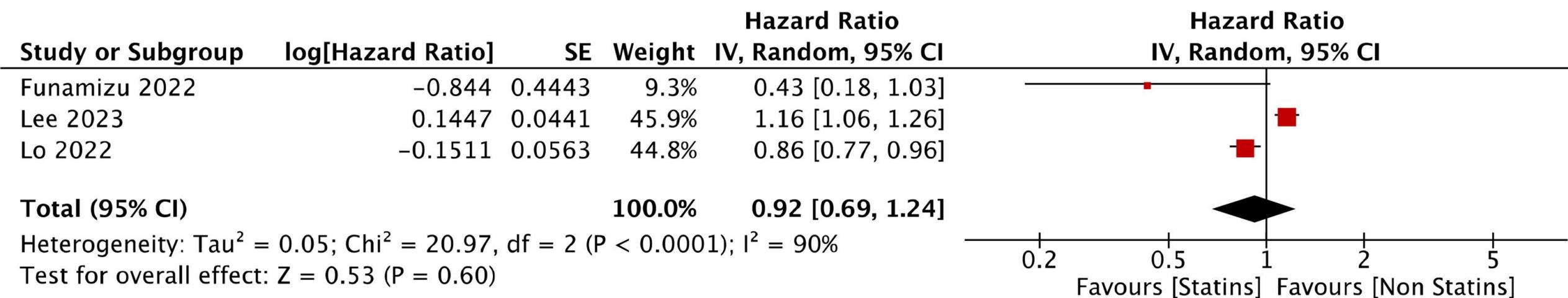
Lee M, Choi WJ, Lee Y, Lee K, Park MW, Myong JP, Kim DW. Association between statin therapy and mortality in patients on dialysis after atherosclerotic cardiovascular diseases. *Sci Rep* 2023;13:10940

Systematic Review and Meta-analysis: PD and Statin: All-Cause Mortality



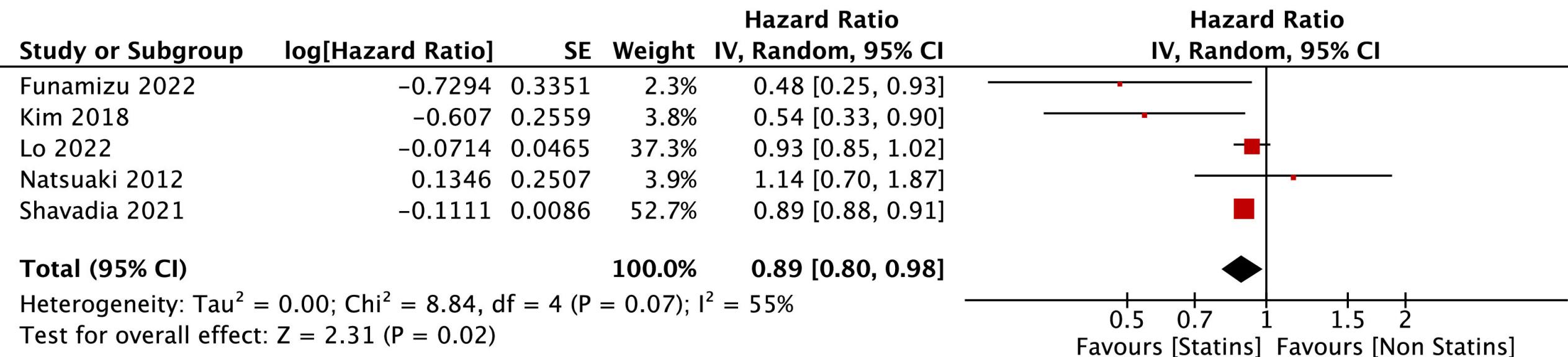
Elkoumi O, Elkoumi A, Elbairy MK, Irfan H, Hageen AW, Beddor A, Khalid N, Mahmoud MAT, Emara A, Alkhazaleh U, Mohamed R, Mohamed A, Elnegiry AK, Nashwan AJ. Impact of statins on mortality and cardiovascular outcomes in dialysis patients with atherosclerotic cardiovascular disease: a systematic review and meta-analysis. Am J Cardiol 2025;256:49-59

Systematic Review and Meta-analysis: PD and Statin: Cardiovascular Mortality



Elkoumi O, Elkoumi A, Elbairy MK, Irfan H, Hageen AW, Beddor A, Khalid N, Mahmoud MAT, Emara A, Alkhazaleh U, Mohamed R, Mohamed A, Elnegiry AK, Nashwan AJ. Impact of statins on mortality and cardiovascular outcomes in dialysis patients with atherosclerotic cardiovascular disease: a systematic review and meta-analysis. Am J Cardiol 2025;256:49-59

Systematic Review and Meta-analysis: PD and Statin: MACE



Elkoumi O, Elkoumi A, Elbairy MK, Irfan H, Hageen AW, Beddor A, Khalid N, Mahmoud MAT, Emara A, Alkhazaleh U, Mohamed R, Mohamed A, Elnegiry AK, Nashwan AJ. Impact of statins on mortality and cardiovascular outcomes in dialysis patients with atherosclerotic cardiovascular disease: a systematic review and meta-analysis. Am J Cardiol 2025;256:49-59



LIPID MANAGEMENT – PRIMARY PREVENTION

Primary prevention of ASCVD

- Supported statin use in observational studies
- Most retrospective studies are propensity-matched

Emulation Target Study of Statin in stage 5 CKD

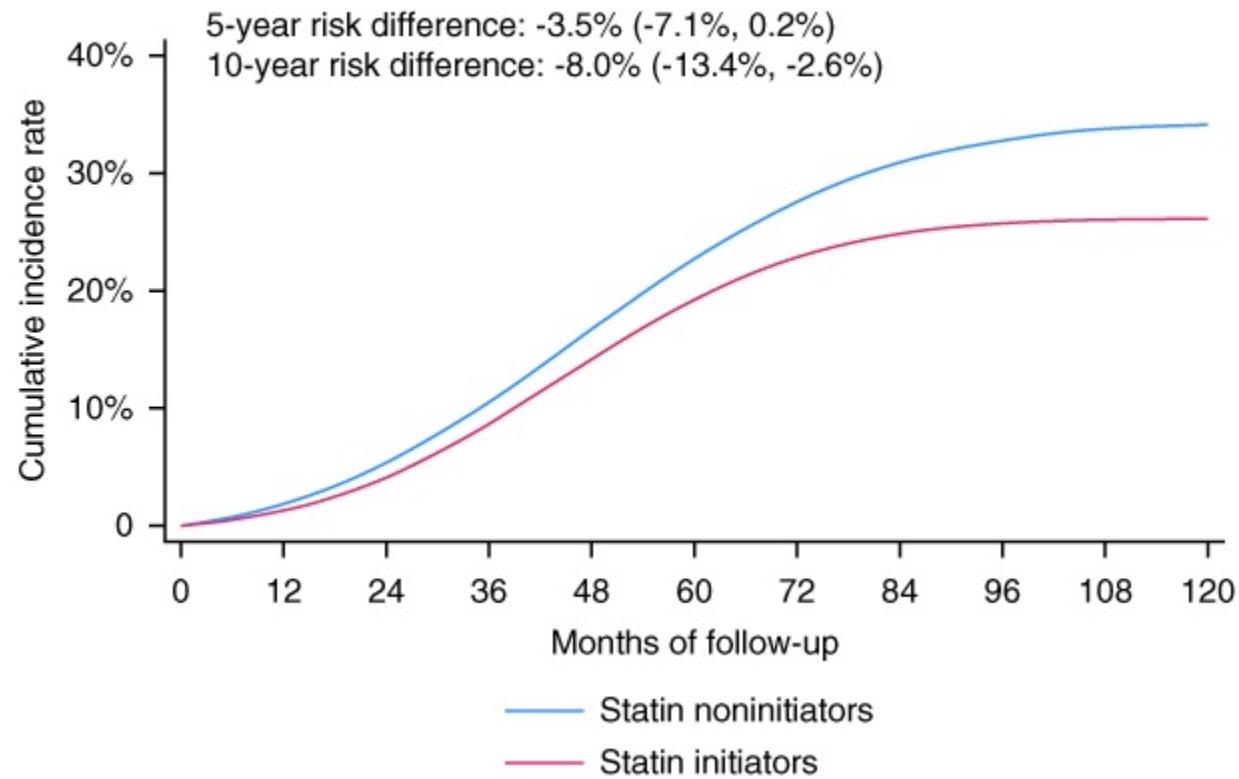
- Electronic health record
- 3019 statin-eligible individuals with kidney failure (eGFR < 15 ml/min/1.73 m²) and elevated LDL cholesterol \geq 100 mg/dl (\geq 2.5 mmol/L)



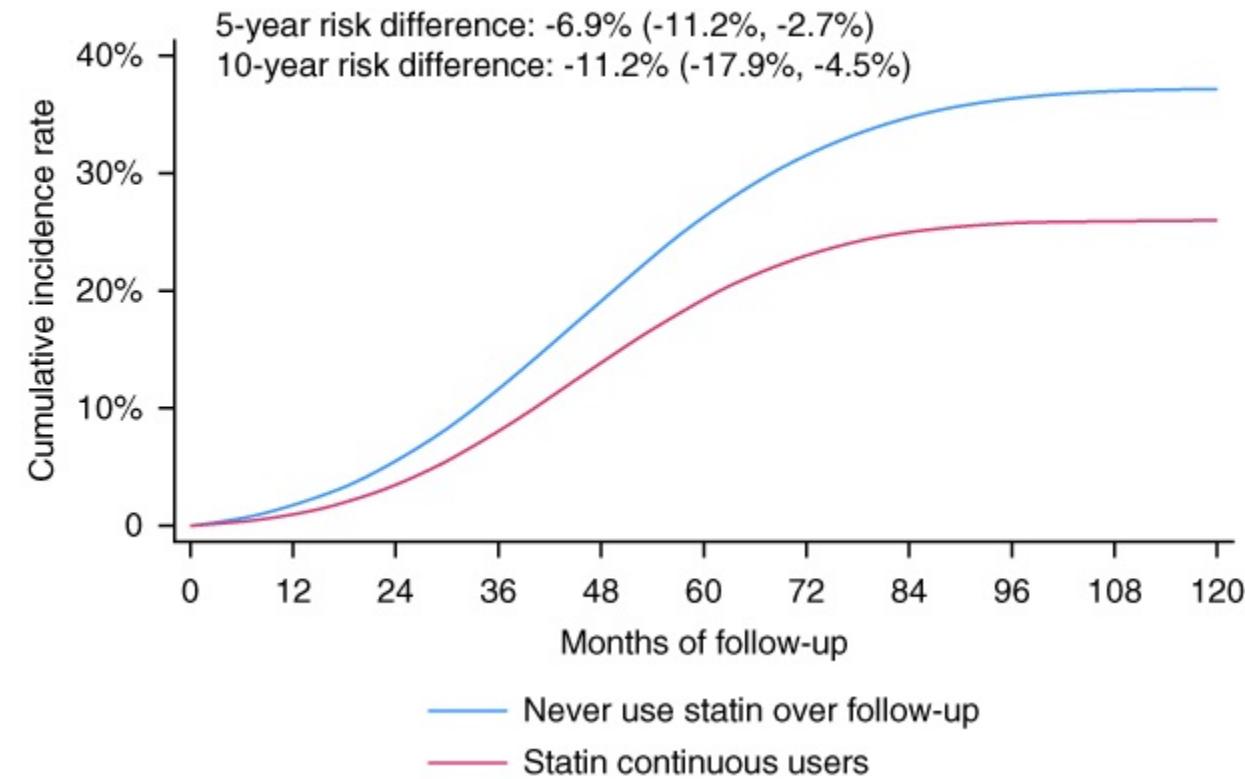
Cheng FW, Xu W, Tang SCW, Wan EYF. Long-term benefits and safety of statins in patients with kidney failure: a target trial emulation study. *J Am Soc Nephrol* 2025;36:882-889

Lower risks of cardiovascular diseases

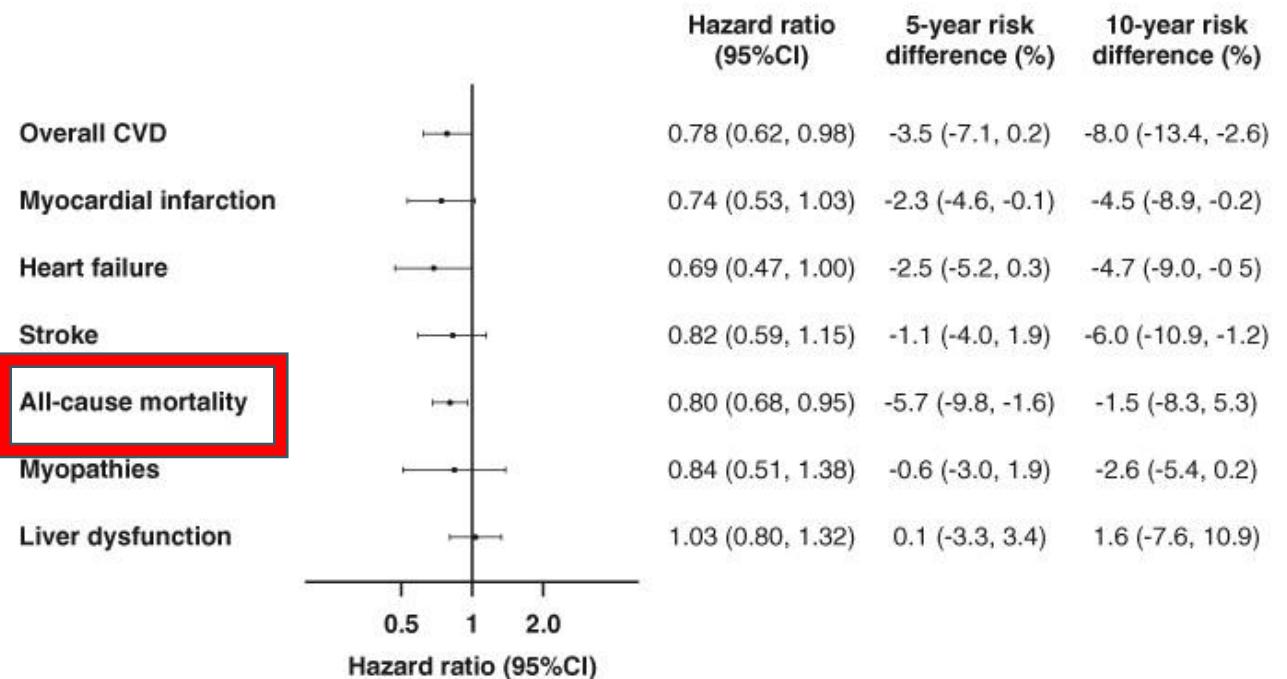
Intention-to-treat analysis



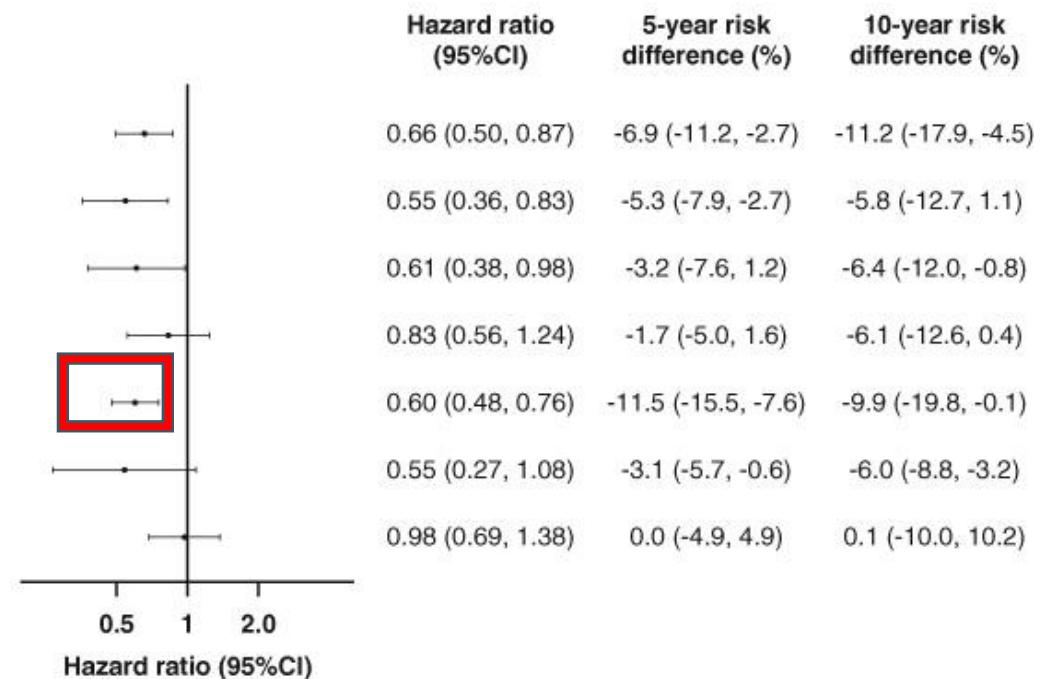
Per-protocol analysis



Intention-to-treat analysis



Per-protocol analysis



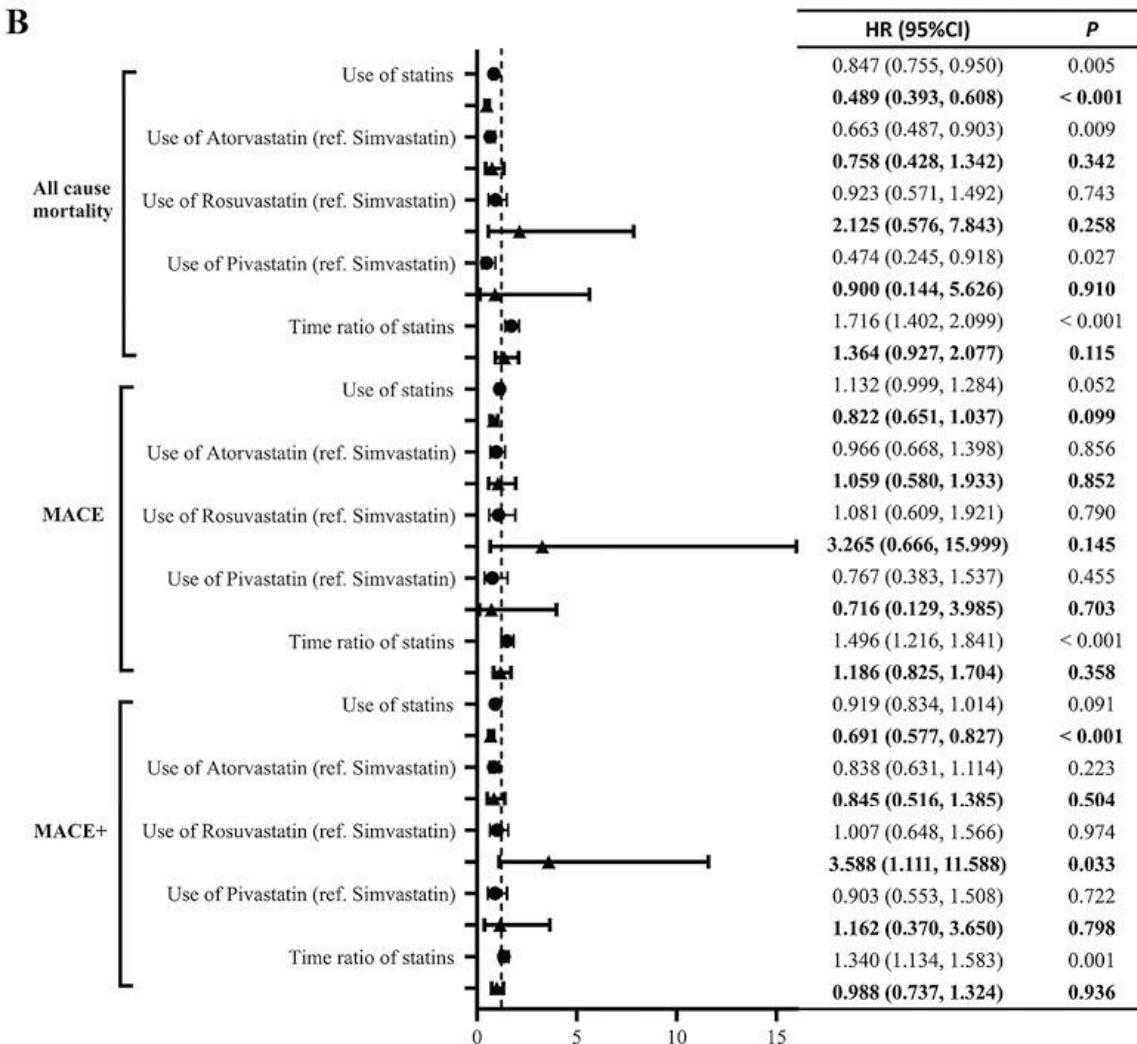
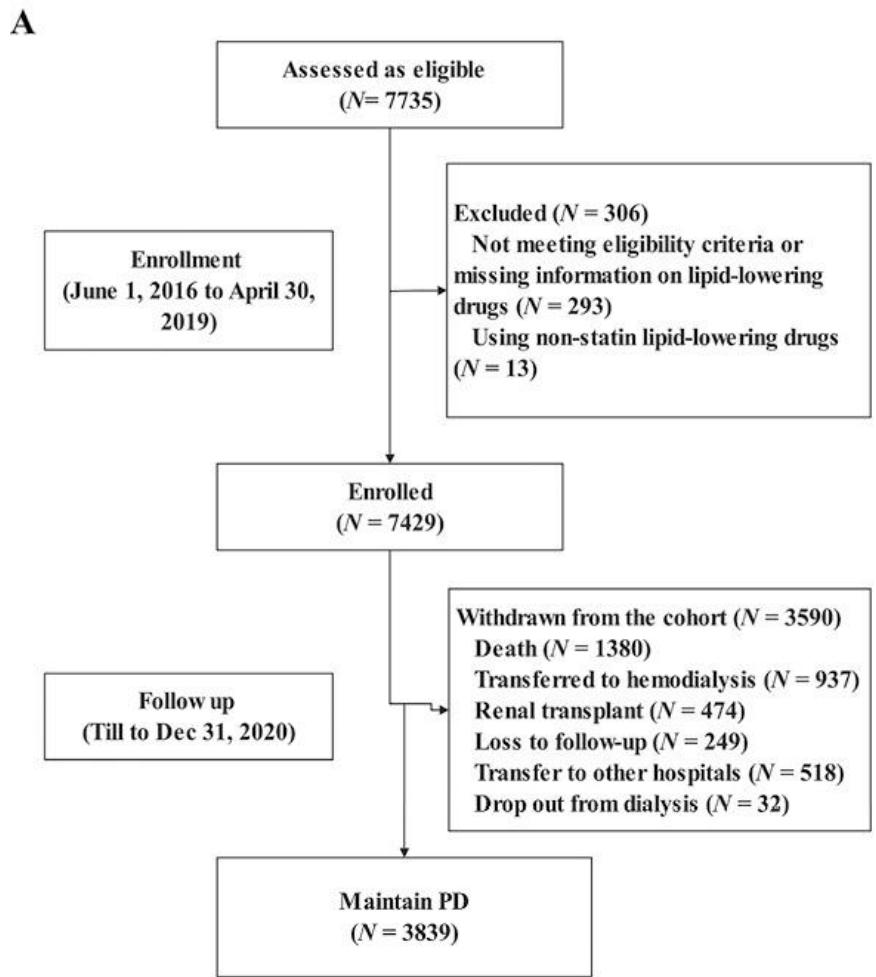
Cheng FW, Xu W, Tang SCW, Wan EYF. Long-term benefits and safety of statins in patients with kidney failure: a target trial emulation study. *J Am Soc Nephrol* 2025;36:882-889

Propensity-matched Cohort in PD

- PD Telemedicine-assisted Platform (PDTAP) Cohort Study, a nationwide large-scale PD cohort study in China
- 7429 patients on PD, from 27 hospitals, followed up for 29.0 months
- 2211 (29.8%) had used statins, median time of medication use 16.7 months

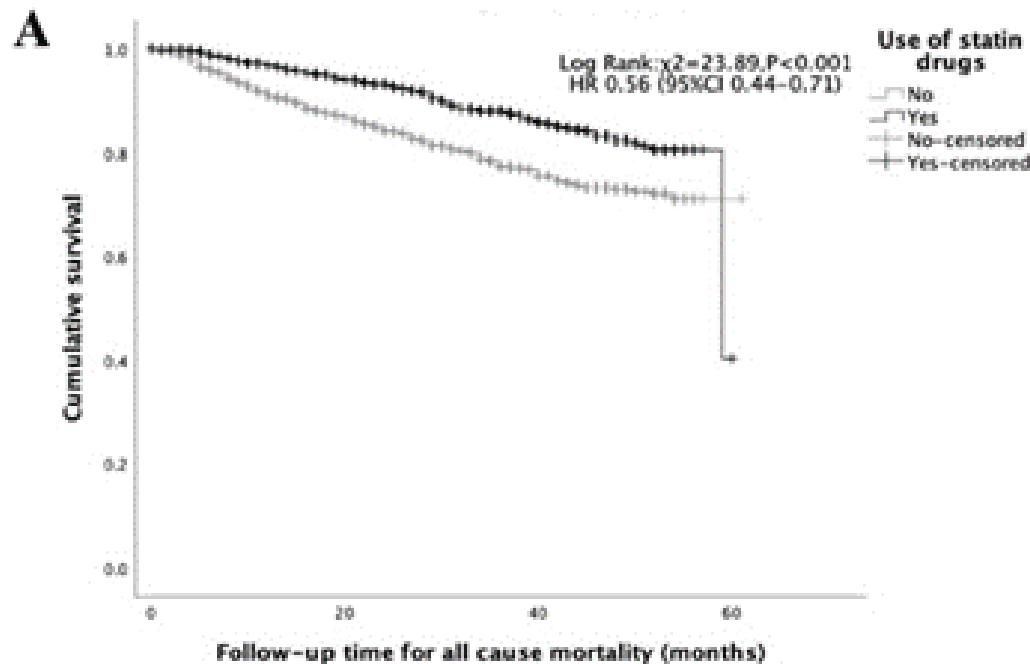


Gao S, Nan L, Li X, et al; PDTAP working group. Associations between statins and all-cause mortality and cardiovascular events among peritoneal dialysis patients: a multi-center large-scale cohort study. Chin Med J (Engl) 2025;138:2856-2858



Gao S, Nan L, Li X, et al; PDTAP working group. Associations between statins and all-cause mortality and cardiovascular events among peritoneal dialysis patients: a multi-center large-scale cohort study. Chin Med J (Engl) 2025;138:2856-2858

Lower All-Cause Mortality with Statin



Gao S, Nan L, Li X, et al; PDTAP working group. Associations between statins and all-cause mortality and cardiovascular events among peritoneal dialysis patients: a multi-center large-scale cohort study. Chin Med J (Engl) 2025;138:2856-2858





LIPID MANAGEMENT

- We suggest considering statin or statin/ezetimibe therapy for PD patients with established atherosclerotic cardiovascular disease (ASCVD) or very high cardiovascular risk (diabetes with hypertension, or low-density lipoprotein cholesterol LDL-C ≥ 190 mg/dL or 4.9 mmol/L), accounting for life expectancy on dialysis and comorbid disease (2B).
- We suggest against routine initiation of statin therapy in patients undergoing PD without ASCVD (2A).

Angiotensin receptor-
neprilysin ARN
inhibitor



HEART FAILURE MANAGEMENT

- We suggest that pharmacological treatment with heart failure indication under guideline-directed medical therapy (GDMT) be optimized in patients receiving PD (Not Graded).



HEART FAILURE MANAGEMENT – RENALISM

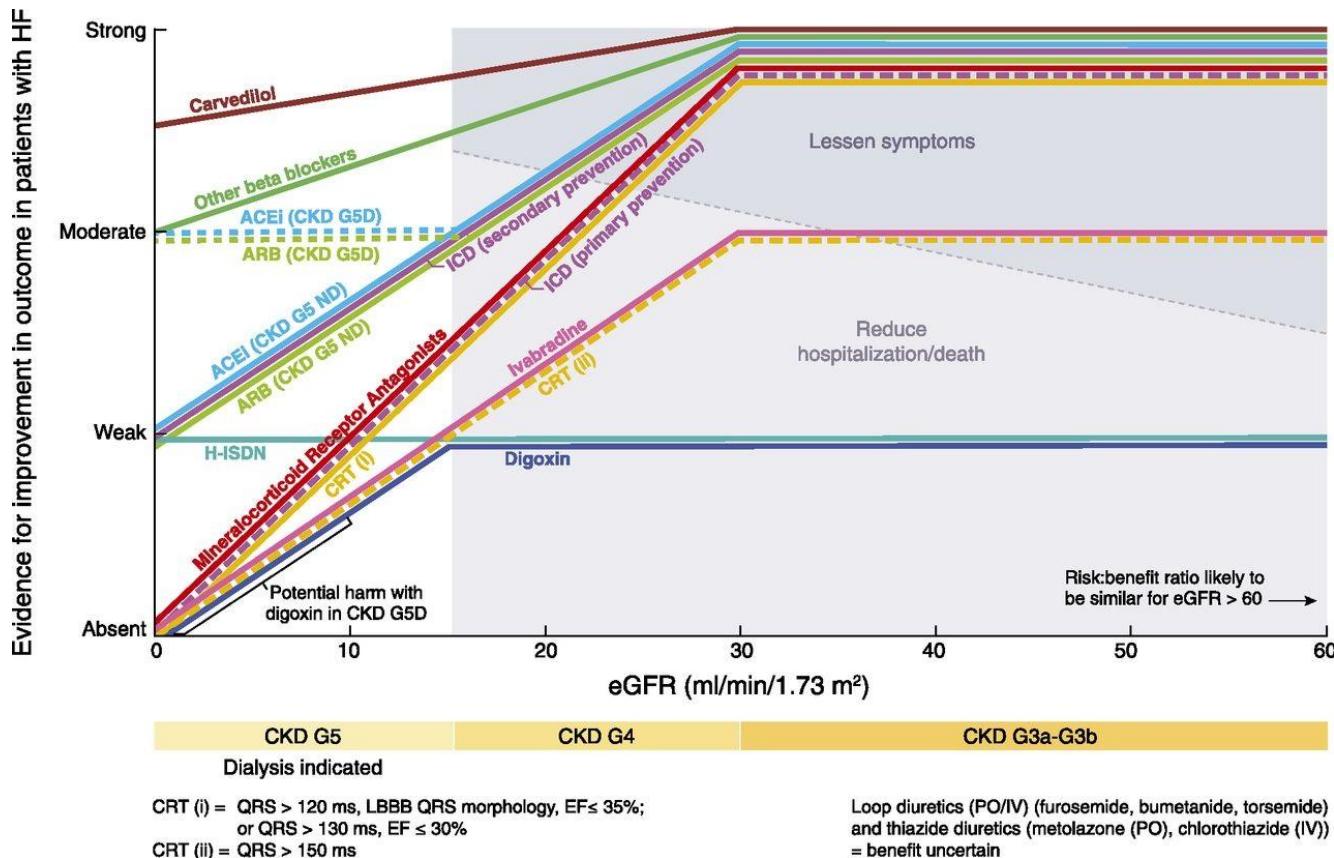
Most landmark RCT for heart failure excluded patients with eGFR < 20

- FINEARTS-HF
- EMPEROR-Reduced
- DAPA-HF
- PARADIGM-HF

Evidence of HFrEF Rx - AJKD Core Curriculum

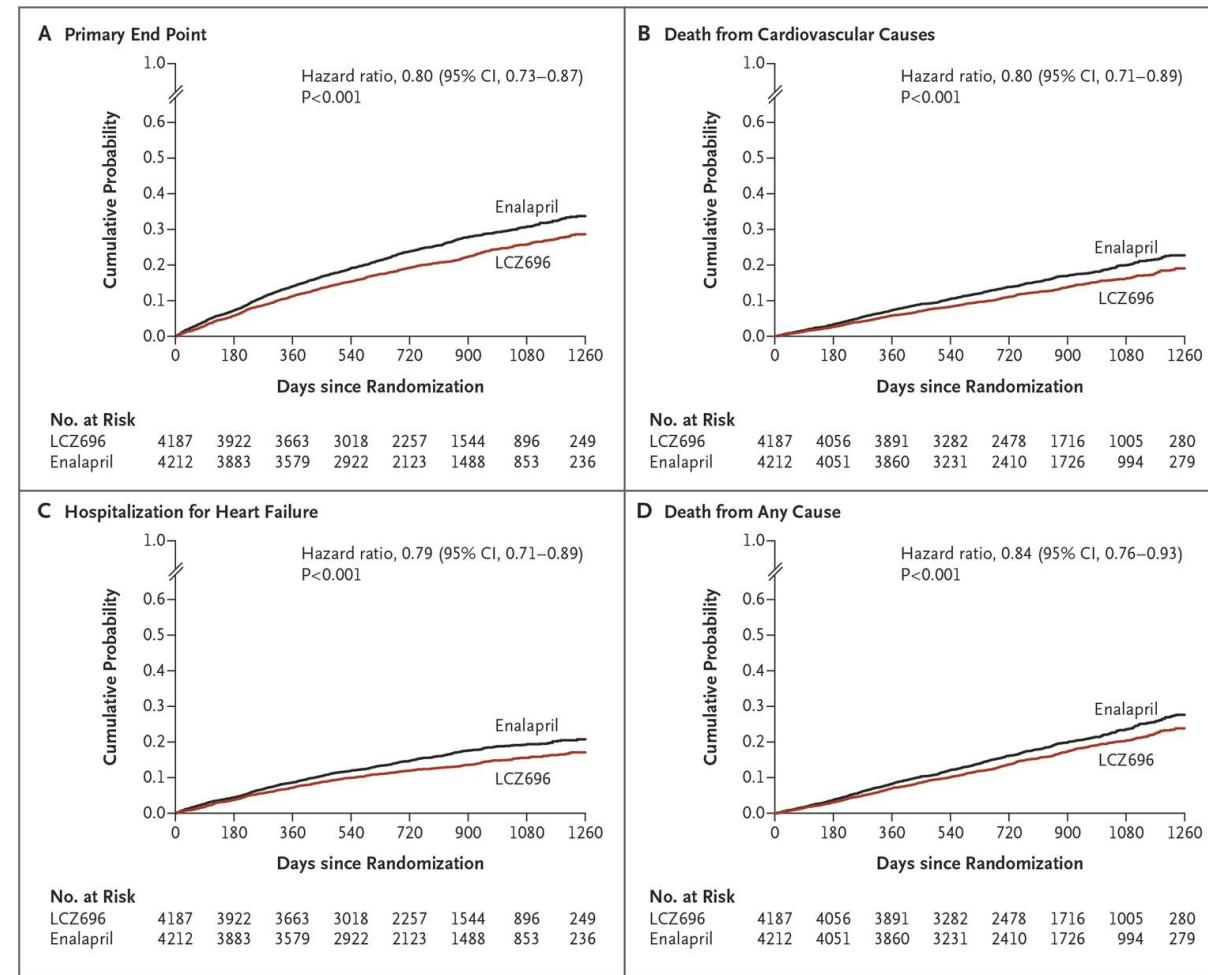
HFrEF therapies	CKD 1 and 2	CKD 3	CKD 4	CKD 5
Beta Blocker	Strong	Strong	Limited	Absent
MRA	Strong	Strong	Limited	Absent
Non-steroidal MRA	Strong	Strong	Strong (up to eGFR > 25 cc/min)	Absent
ARNi	Strong	Strong	Limited	Absent
ACEi/ARB	Strong	Strong	Limited	Absent
Diuretics	Absent	Absent	Absent	Absent
SGLT2i	Strong	Strong	Strong (eGFR > 20 cc/min)	Limited

KDIGO Consensus Conference Report



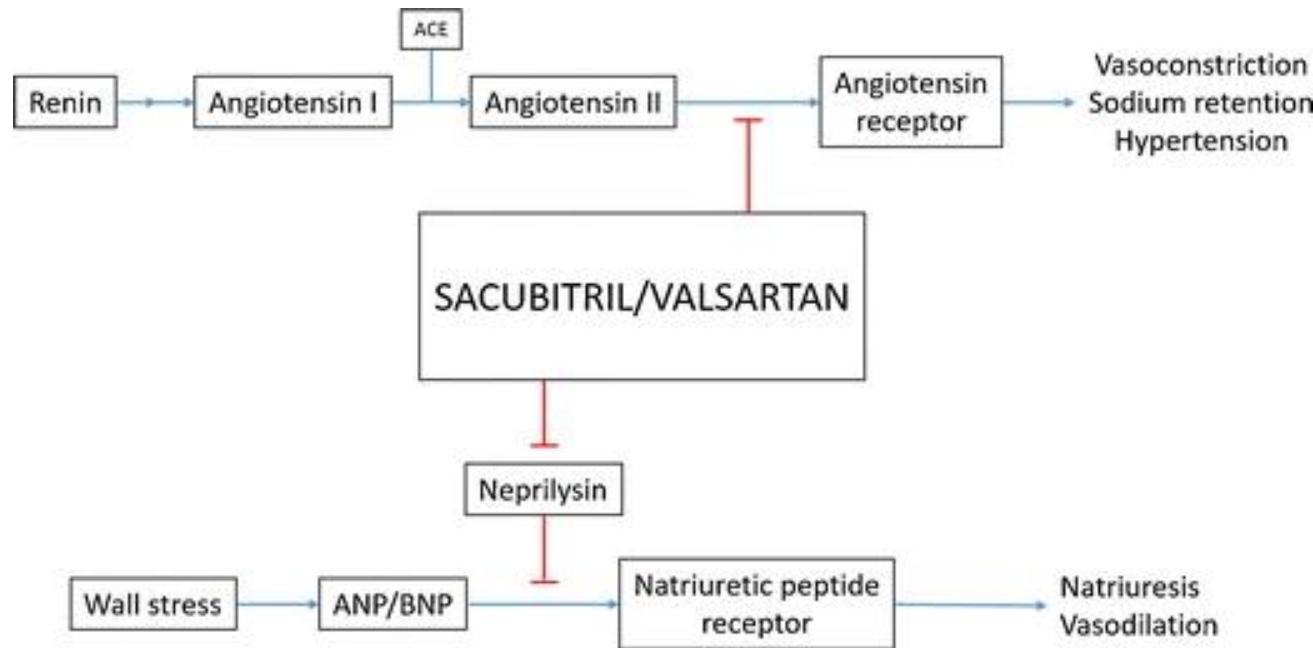
Banerjee D, Rosano G, Herzog CA. Management of heart failure patient with CKD. Clin J Am Soc Nephrol 2021;16:1131-1139

PARADIGM-HF – ARNI



McMurray JJ, Packer M, Desai AS, Gong J, Lefkowitz MP, Rizkala AR, Rouleau JL, Shi VC, Solomon SD, Swedberg K, Zile MR; PARADIGM-HF Investigators and Committees. Angiotensin-neprilysin inhibition versus enalapril in heart failure. *N Engl J Med* 2014;371:993-1004

SACUBITRIL/VALSARTAN



Kang G, Banerjee D. Neprilysin inhibitors in cardiovascular disease. *Curr Cardiol Rep* 2017;19:16

Observational Data

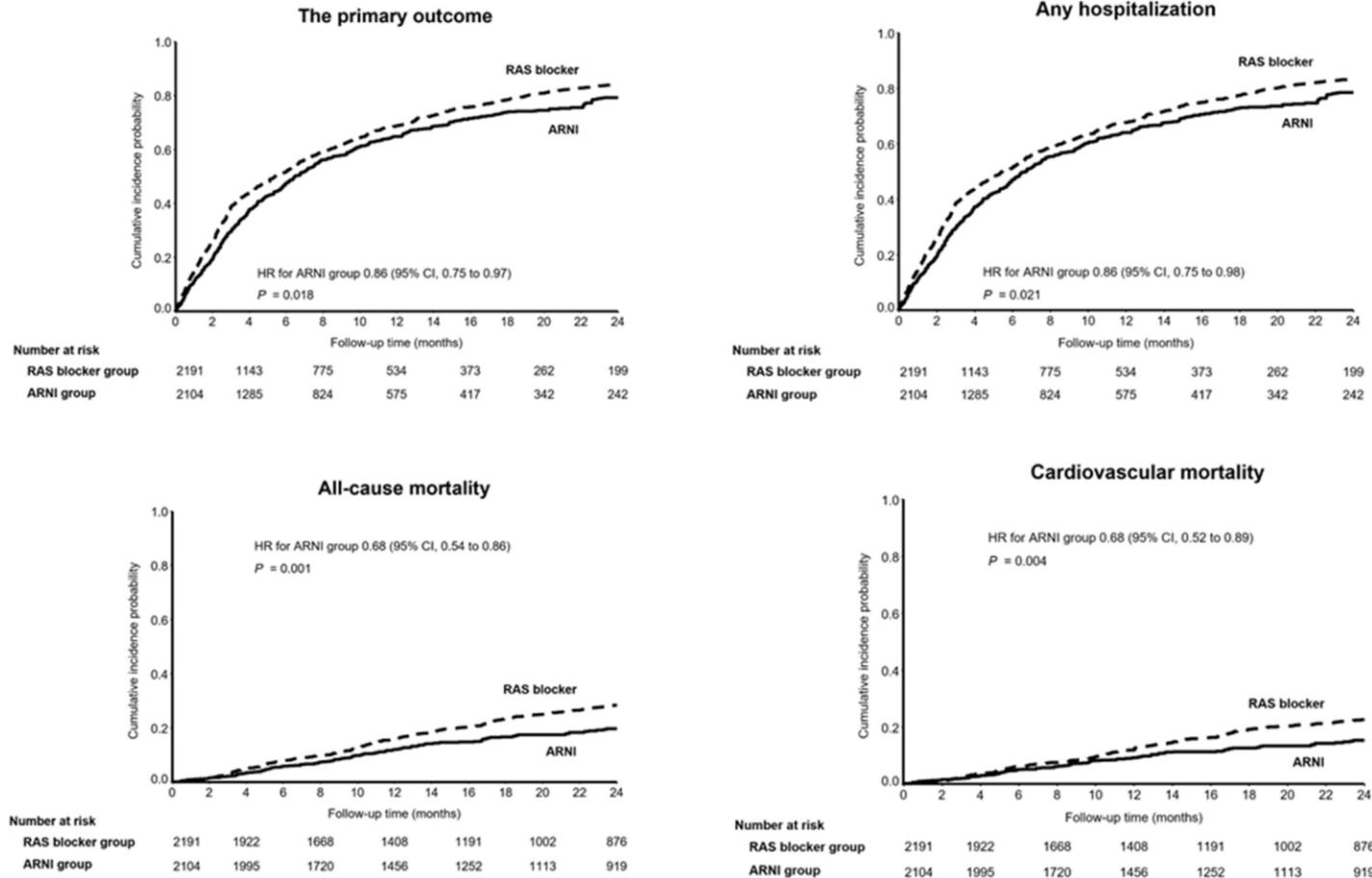
National Health Insurance Service database

- Patients with HFrEF and kidney failure receiving dialysis who were prescribed either ARNI or renin-angiotensin system (RAASi)
- Inverse probability of treatment weighting: 2104 patients on ARNI and 2191 on RAASI



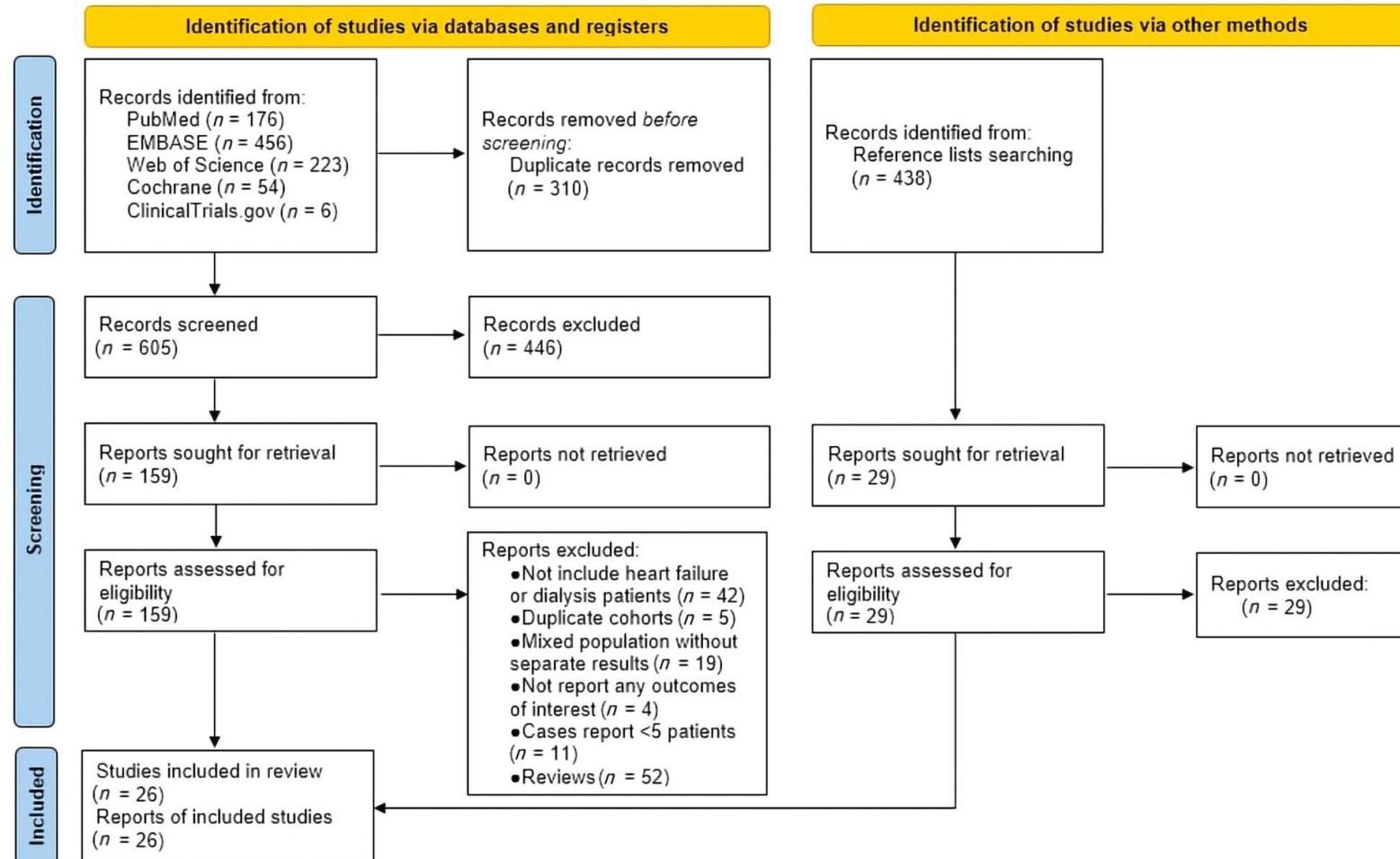
Jung MH, Cho DH, Choi J, Kim MN, Lee CJ, Son JW, Kim Y, Youn JC, Yoo BS.
Angiotensin receptor-neprilysin inhibitors in concurrent heart failure with
reduced ejection fraction and kidney failure. ESC Heart Fail 2025;12:3405-3415

Lower all-cause mortality, any hospitalization and cardiovascular mortality with ARNI

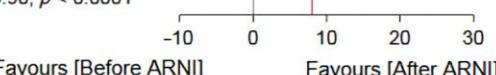
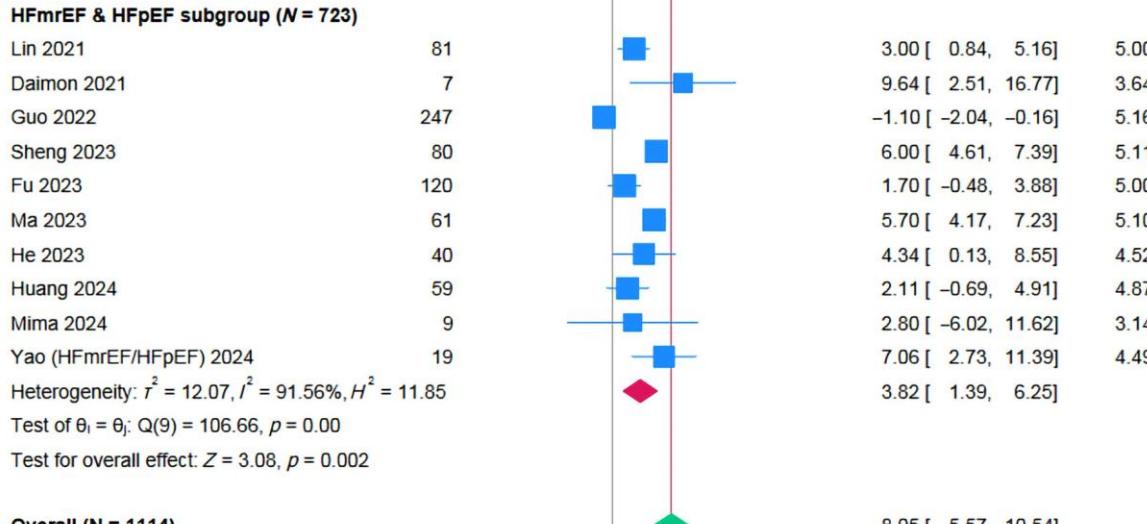
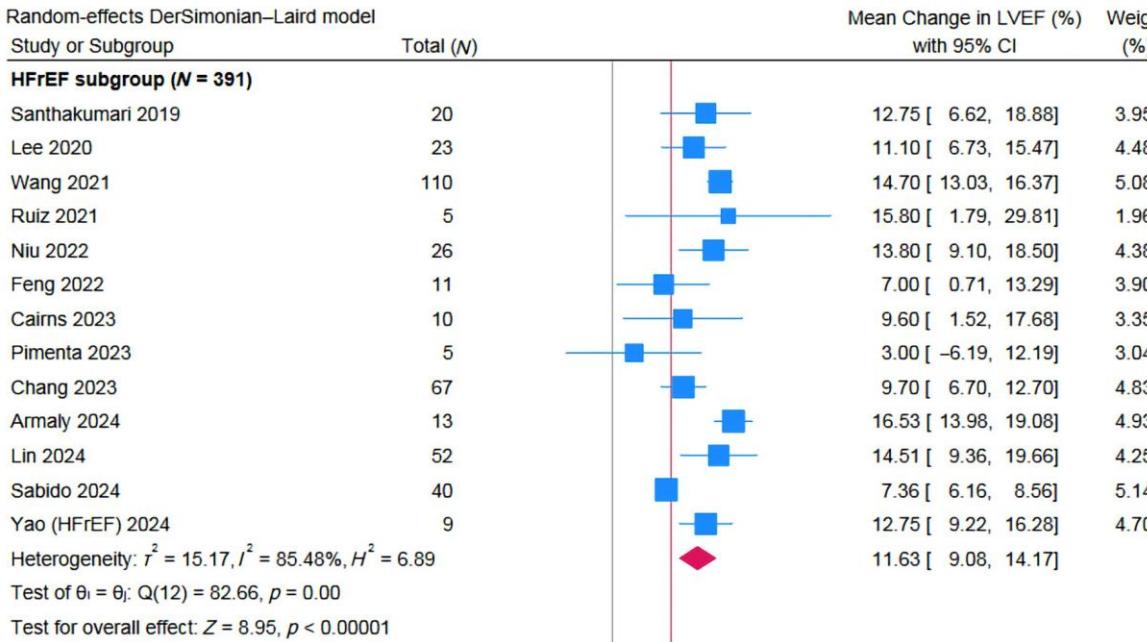


Jung MH, Cho DH, Choi J, Kim MN, Lee CJ, Son JW, Kim Y, Youn JC, Yoo BS. Angiotensin receptor-neprilysin inhibitors in concurrent heart failure with reduced ejection fraction and kidney failure. ESC Heart Fail 2025;12:3405-3415

ARNI



Nguyen DV, Le TN, Truong BQ, Nguyen HTT. Efficacy and safety of angiotensin receptor-neprilysin inhibition in heart failure patients with end-stage kidney disease on maintenance dialysis: a systematic review and meta-analysis. Eur J Heart Fail 2025;27:72-84



Improvement in left ventricular ejection fraction (LVEF)

Nguyen DV, Le TN, Truong BQ, Nguyen HTT. Efficacy and safety of angiotensin receptor-neprilysin inhibition in heart failure patients with end-stage kidney disease on maintenance dialysis: a systematic review and meta-analysis.

Eur J Heart Fail 2025;27:72-84

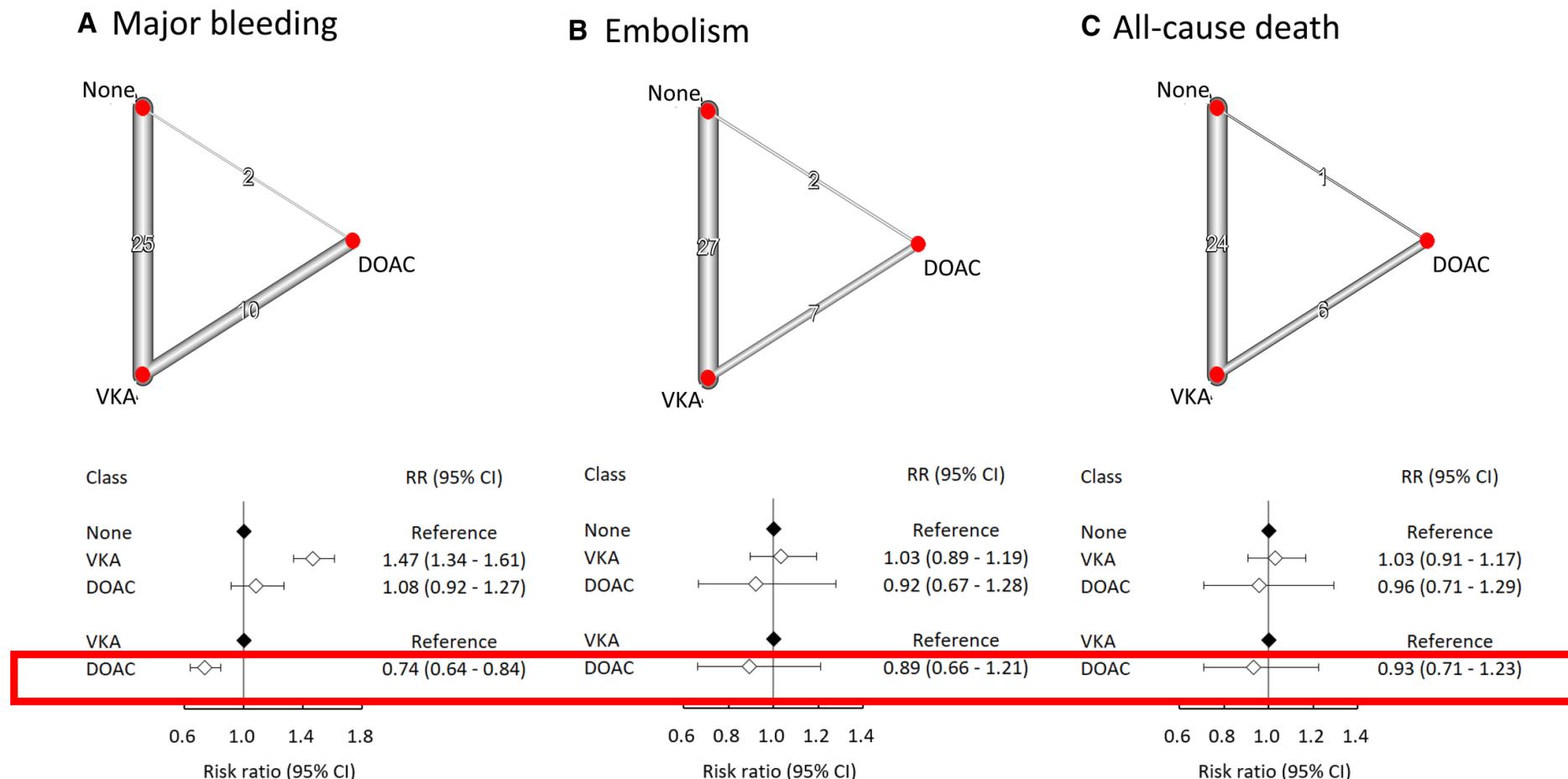


HEART FAILURE MANAGEMENT

- We recommend sacubitril-valsartan use in patients on PD with heart failure with a reduced ejection fraction (HFrEF) and a history of New York Heart Association (NYHA) class II to III symptoms; ACEi and ARB should be considered in patients with contraindications, intolerance, or inaccessibility to ARNI (1C).

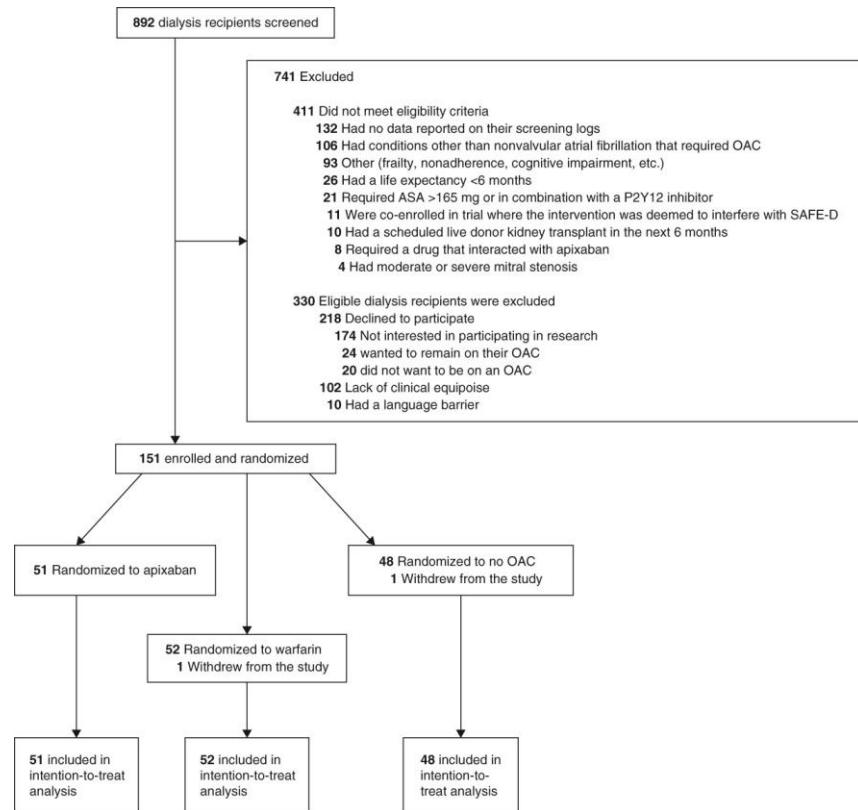
ANTICOAGULATION

Systematic Review and Meta-analysis



RCT: SAFE-D

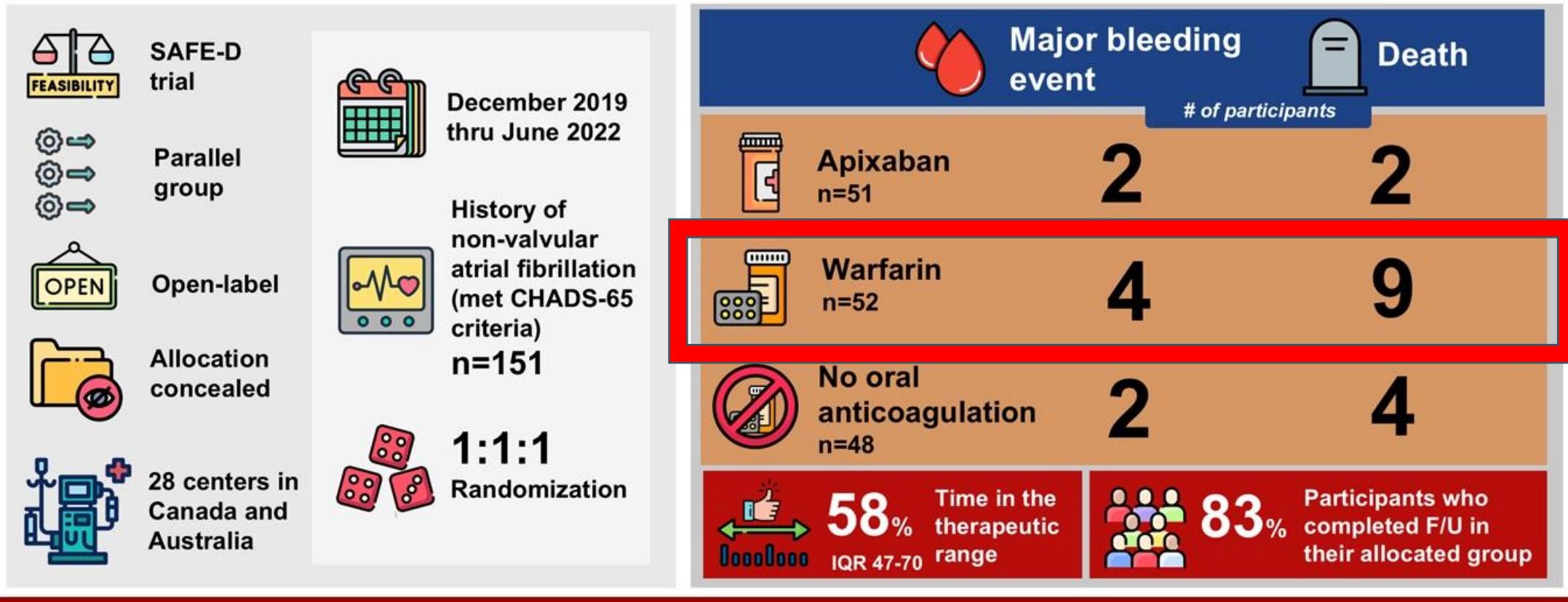
Strategies for the Management of Atrial Fibrillation in Patients Receiving Dialysis (SAFE-D)



Harel Z, Smyth B, Badve SV, Blum D, Beaubien-Souigny W, Silver SA, Clark E, Suri R, Mavrakanas TA, Sasal J, Prasad B, Eikelboom J, Tennankore K, Rigatto C, Prce I, Madore F, Mac-Way F, Steele A, Zeng Y, Sholzberg M, Dorian P, Yan AT, Sood MM, Gladstone DJ, Tseng E, Kitchlu A, Walsh M, Sapir D, Oliver MJ, Krishnan M, Kiaii M, Wong N, Kotwal S, Battistella M, Acedillo R, Lok C, Weir M, Wald R. Anticoagulation for patients with atrial fibrillation receiving dialysis: a pilot randomized controlled trial. *J Am Soc Nephrol* 2025;36:901-910



Strategies for the Management of Atrial Fibrillation in patients Receiving Dialysis (SAFE-D)



Conclusions: This study demonstrated the feasibility of recruitment and adherence in a trial that compared different anticoagulation strategies in patients with atrial fibrillation receiving dialysis.

Ziv Harel, Brendan Smyth, Sunil V. Badve, et al. *Anticoagulation for Patients with Atrial Fibrillation Receiving Dialysis: A Pilot Randomized Controlled Trial*. JASN doi: 10.1681/ASN.0000000000000495.
Visual Abstract by Edgar Lerma, MD, FASN

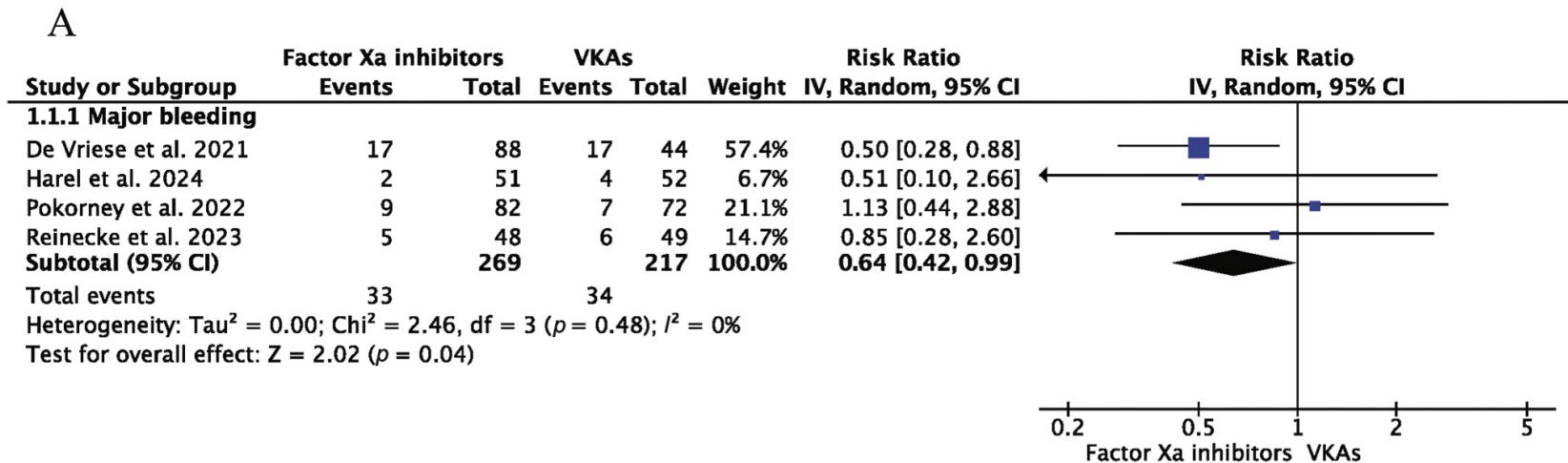
Systematic Review and Meta-analysis: RCT

Studies	Study type	Trial acronyms	Location	Comparison	Dialysis characteristics	Sample size	Age (years)	Female (%)	DOAC type and dosage	TTR for VKA group	Follow-up
Reinecke et al. (2023)	PROBE RCT	AXADIA-AFNET 8 trial; NCT02933697	Germany; 2017–2022	Apixaban versus Phenprocoumon	Hemodialysis: 93.8%; Peritoneal: 6.2%	n = 97 API (48) and phenprocoumon (49)	74.7	30%	API; 2.5 mg BID	50.7%	API: 429 days VKA: 506 days
Pokorney et al. (2022)	Multicenter RCT; PROBE	RENAL-AF trial; NCT02942407	42 U.S. clinical sites; 2017–2019	API versus Warfarin	Hemodialysis	n = 154 API (82) and warfarin (72)	68.0	56%	API; 5 mg BID (2.5 mg BID for patients ≥80 years, weight ≤60 kg)	44%	API: 330 days VKA: 340 days
De Vries et al. (2021)	Multicenter RCT	Valkyrie trial; NCT03799822	3 sites in Belgium; 2015–2019	RIV versus VKA versus RIV + Vitamin K2	Hemodialysis	n = 132 Rivaroxaban (88) and VKA (44)	RIV: 79.9 and VKA: 80.3	33%	RIV; 10 mg QD	48%	1.88 years
Harel et al. (2024)	Pilot RCT	SAFE-D trial; NCT03987711	28 centers in Canada and Australia; 2019–2022	API versus Warfarin versus No OAC	Hemodialysis: 93%; Peritoneal: 7%	n = 103 API (51) and warfarin (52)	API: 72 and VKA: 71	22.3%	API; 5 mg BID (2.5 mg BID for patients ≥80 years, weight ≤60 kg)	58%	26 weeks

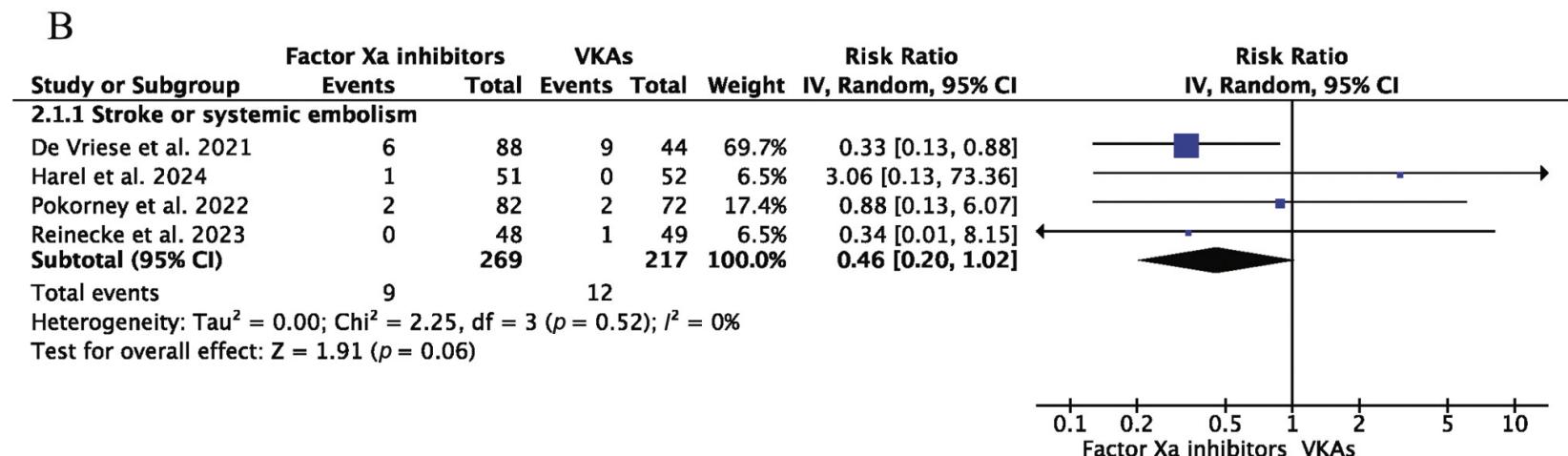
Kaisaier W, Chen Y, Lip GYH, Liu C, Zhu W. Safety and efficacy of factor Xa inhibitors in atrial fibrillation patients on dialysis: evidence from four randomized controlled trials. *Thromb Haemost* 2025 Epub ahead of print

Systematic Review and Meta-analysis: RCT

Primary safety



Primary efficacy



Factor Xa Inhibitors vs. Vitamin K Antagonists in Atrial Fibrillation Patients on Dialysis



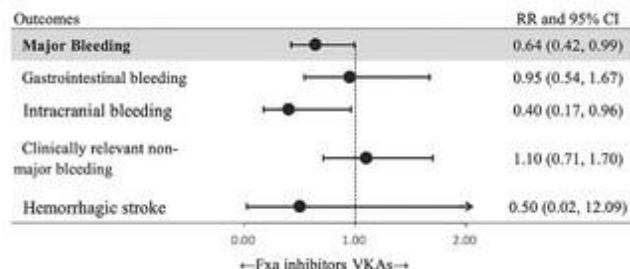
486 dialysis-dependent
AF patients



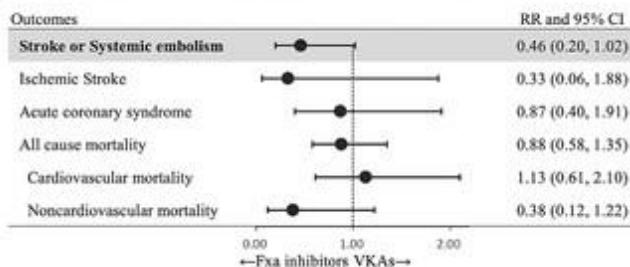
4 randomized
controlled trials



Safety



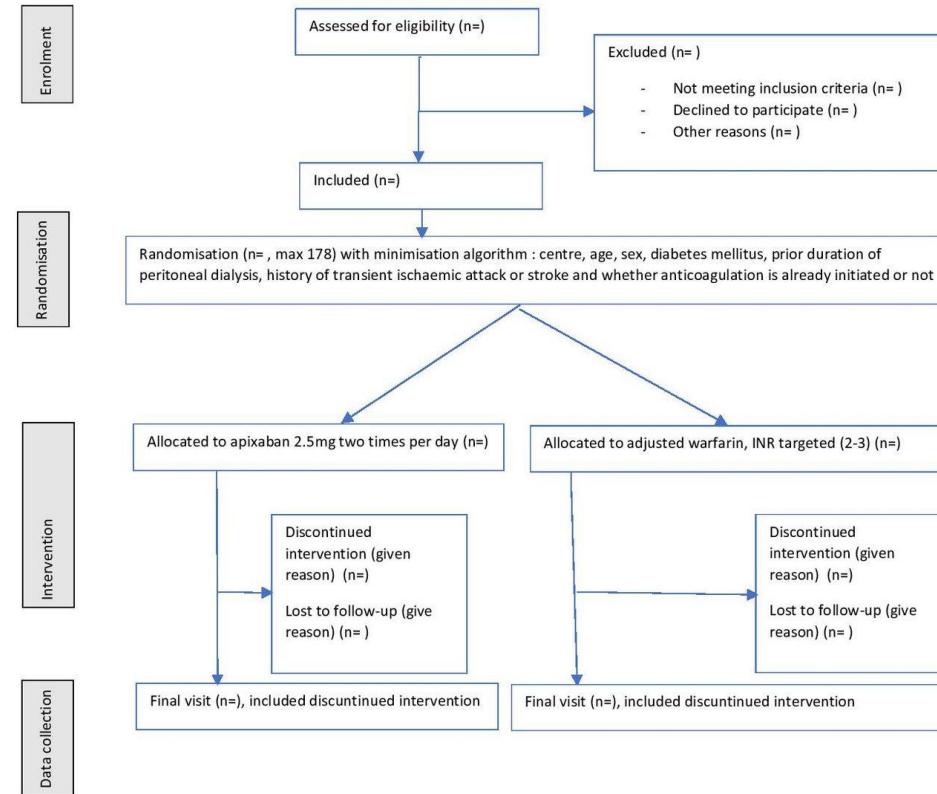
Efficacy



Conclusion: Factor Xa inhibitors may offer a safer alternative to VKAs for AF patients on dialysis, with a lower risk of bleeding and similar risks of stroke and mortality.

Visual summary. Summary for effect estimates of factor Xa inhibitors versus VKAs in atrial fibrillation patients on dialysis, expressed as RR and 95% CI. AF, atrial fibrillation; CI, confidence interval; IV, inverse variance; RR, risk ratio; VKA, vitamin K antagonist.

RCT Study in PD – to be finished 2027



Planned sample size:
178 participants from 20 PD centres

Ficheux M, Peyro-Saint-Paul L, Balayn D, Lecrux B, Brossier M, Morin A, Lanot A, Peron C, Boulanger M, Brionne M, Beygui F, Parienti JJ, Lobbedez T, Béchade C. Safety and efficacy of apixaban versus warfarin in peritoneal dialysis patients with non-valvular atrial fibrillation: protocol for a prospective, randomised, open-label, blinded endpoint trial (APIDP2). *BMJ Open* 2024;14:e089353



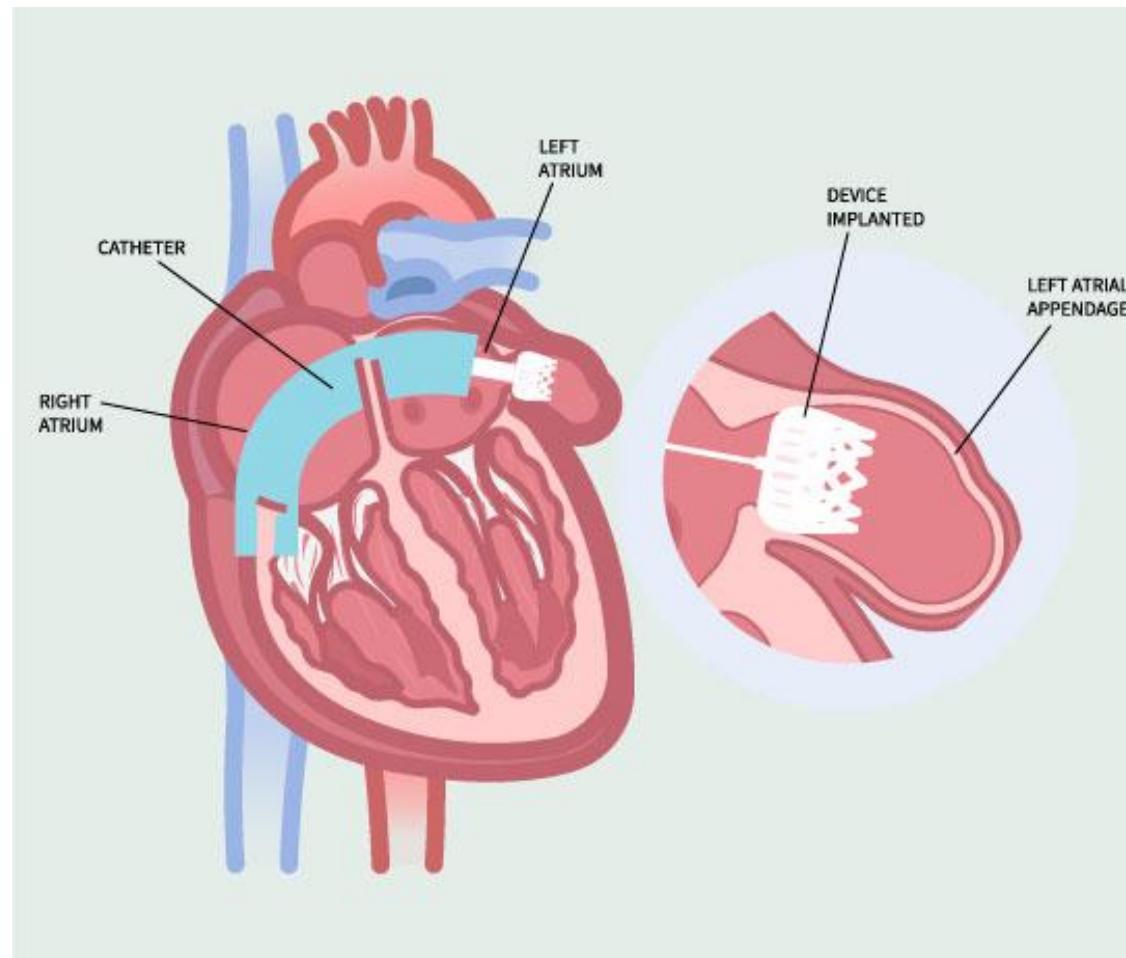
CARDIAC ARRHYTHMIAS MANAGEMENT

- We suggest shared medical decision-making to **consider direct oral anticoagulation (DOAC) in high-risk patients** on PD with atrial fibrillation **(2C)**.
- We suggest no anticoagulation instead of a vitamin K antagonist in patients on PD with a low risk of stroke or high bleeding risk (2D).

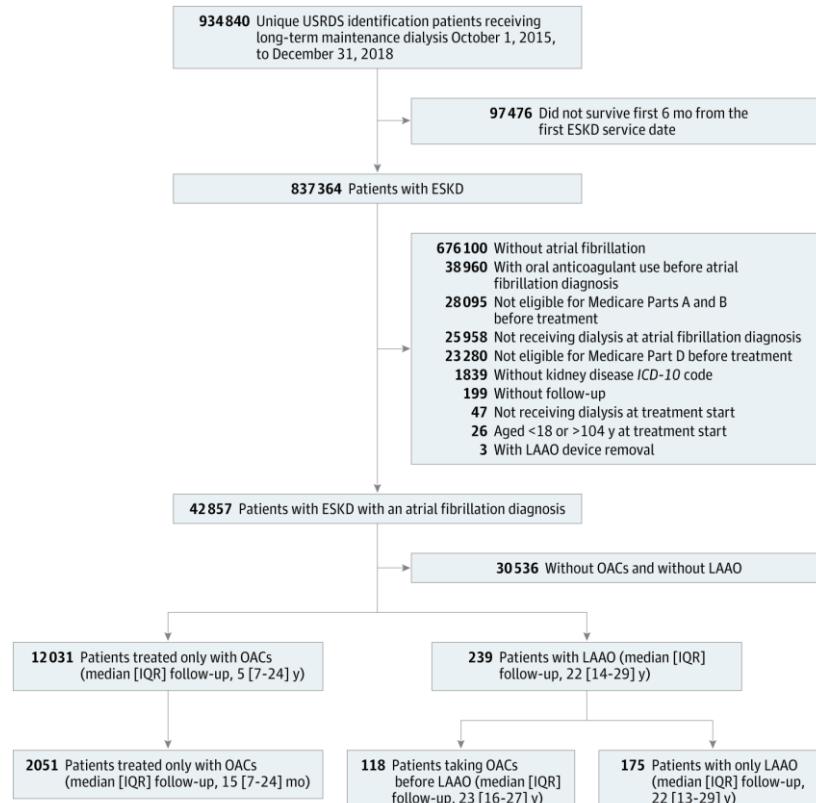
LAAO



LEFT ATRIAL APPENDAGE OCCLUSION



LAAO – Propensity matching



USRDS data with Medicare claims

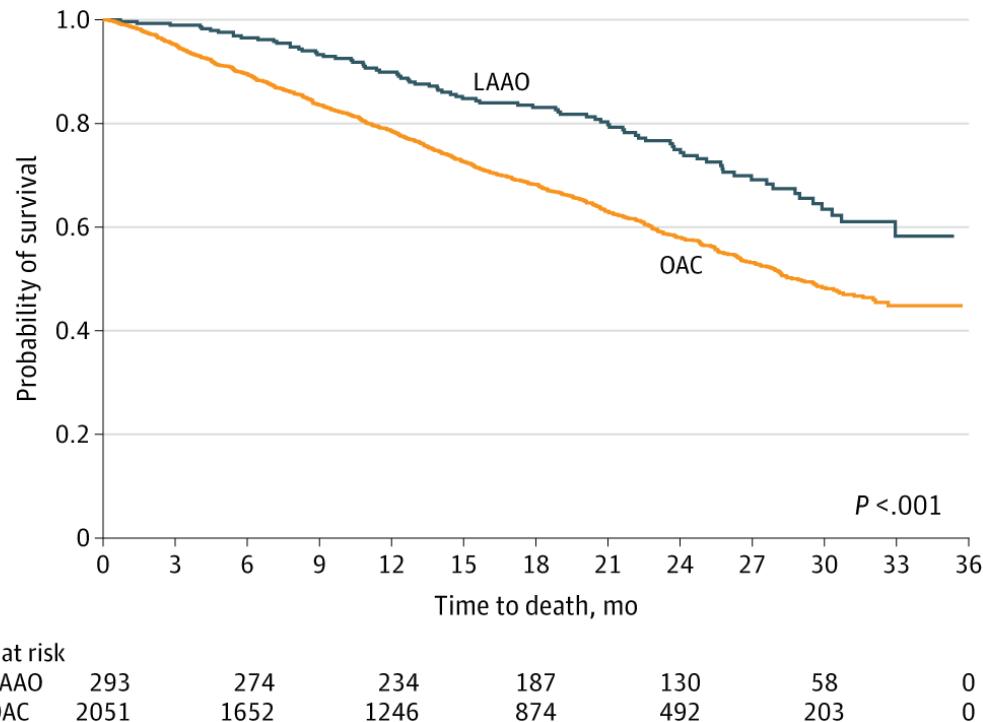
N = 2344 patients on dialysis (150 on PD, 6.4%)

Two groups with propensity score matching
LAAO, oral anticoagulation



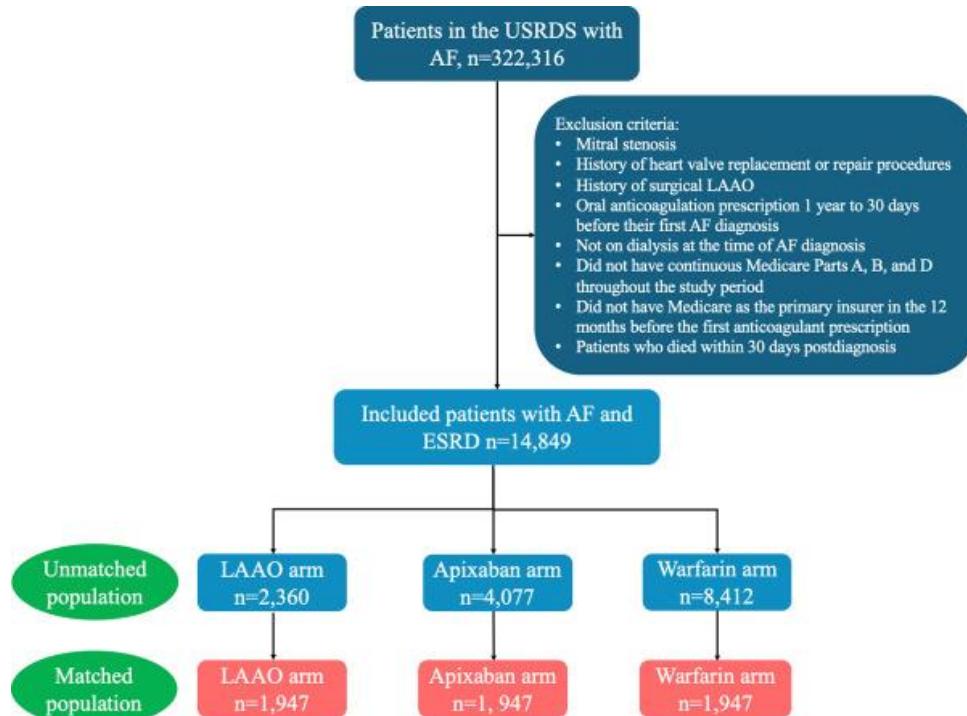
Dhar G, Phadnis MA, Hunt SL, Du HE, Ong V, Khandekar N, Shireman TI, Lynch D, Randhawa S, Deshmukh A, Vallurapalli S, Jain N. Left atrial appendage occlusion vs anticoagulants in dialysis with atrial fibrillation. JAMA Netw Open 2025;8:e2530990

LAAO – Lower all-cause mortality



Dhar G, Phadnis MA, Hunt SL, Du HE, Ong V, Khandekar N, Shireman TI, Lynch D, Randhawa S, Deshmukh A, Vallurapalli S, Jain N. Left atrial appendage occlusion vs anticoagulants in dialysis with atrial fibrillation. JAMA Netw Open 2025;8:e2530990

LAAO – Propensity matching



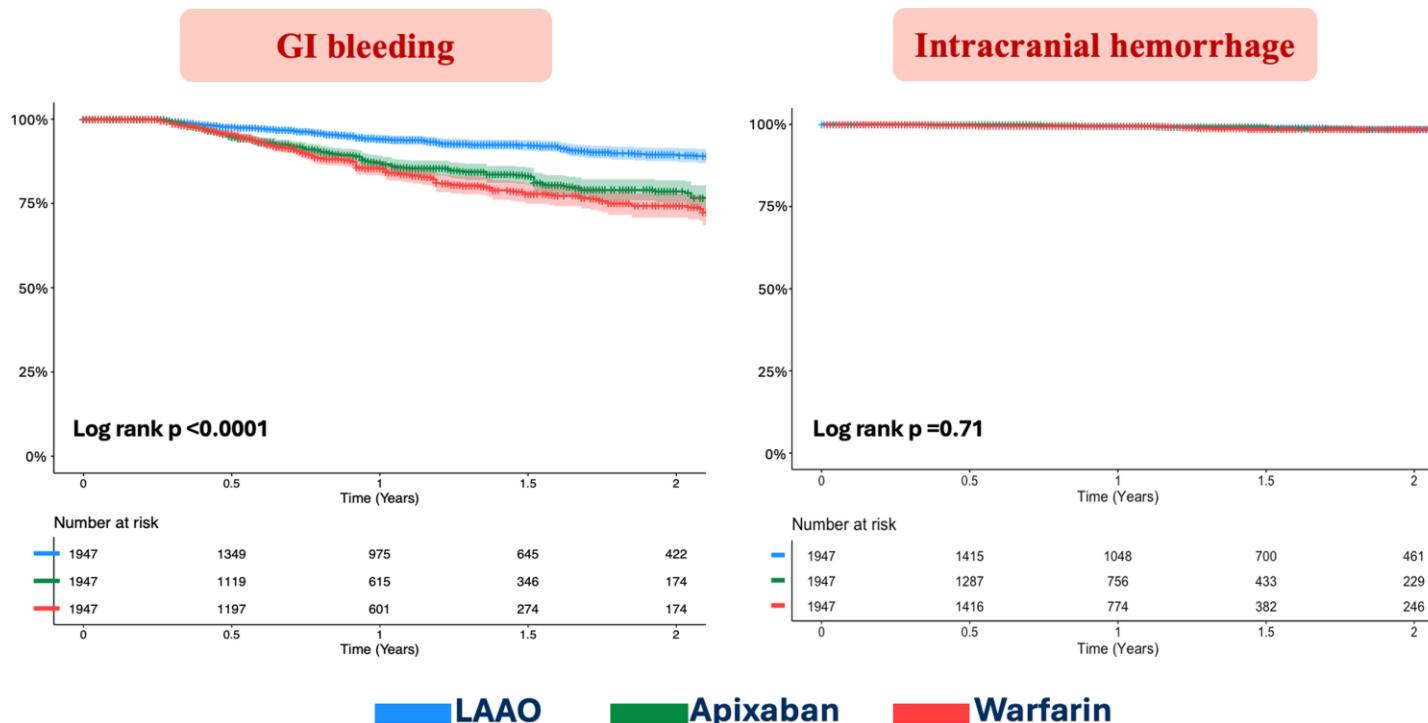
14,849 patients with kidney failure and AF

Three groups with propensity score matching
LAAO, apixaban, warfarin



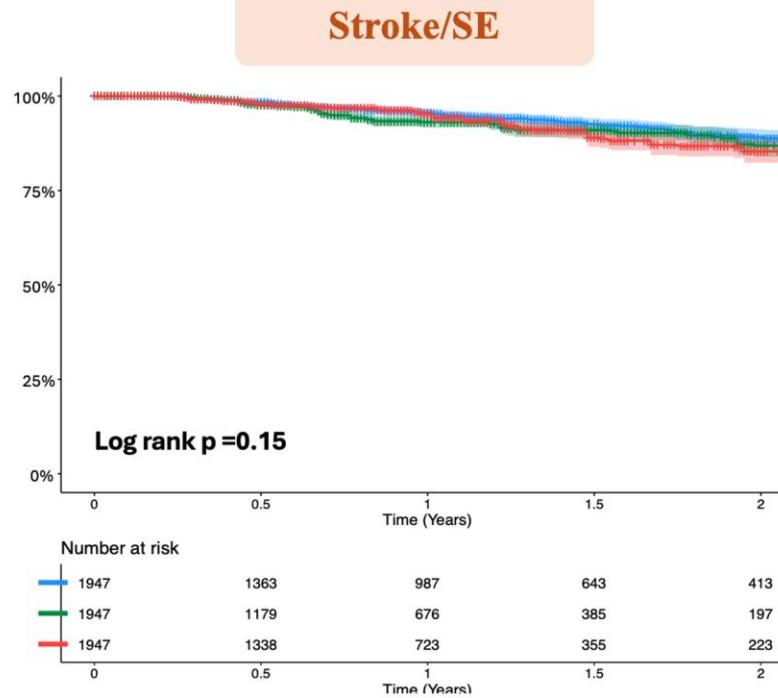
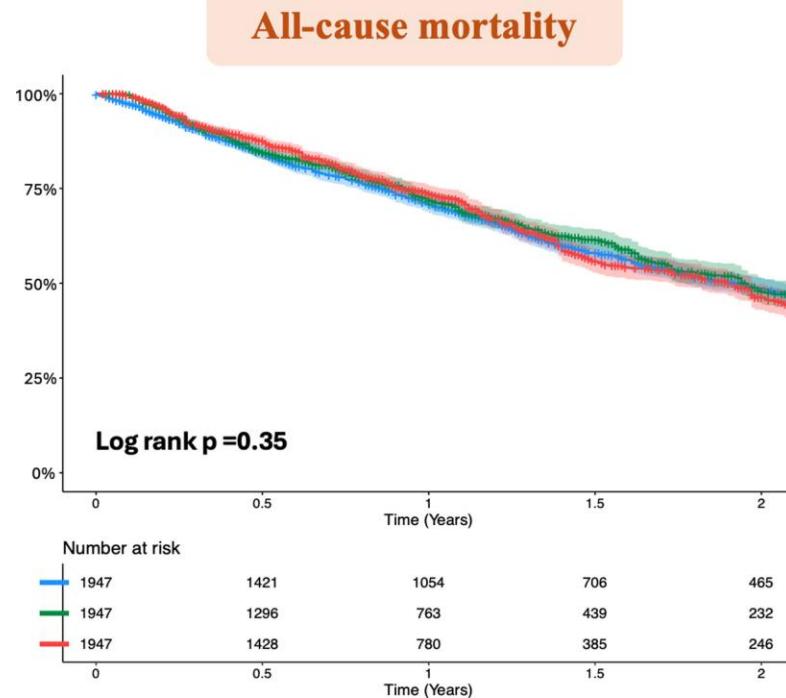
Al-Abcha A, Saleh G, Sledge H, Abbasi M, Ismayl M, Killu AM, Simard T, Liu X, Hibbert B, Friedman PA, Siontis KC, Alkhouri M. Outcomes of stroke prevention strategies in patients with atrial fibrillation and end-stage renal disease. JACC Cardiovasc Interv 2025;18:2614-2624

LAAO: Lowest major and GI bleeding



Al-Abcha A, Saleh G, Sledge H, Abbasi M, Ismayl M, Killu AM, Simard T, Liu X, Hibbert B, Friedman PA, Siontis KC, Alkhouri M. Outcomes of stroke prevention strategies in patients with atrial fibrillation and end-stage renal disease. JACC Cardiovasc Interv 2025;18:2614-2624

LAAO: Lowest primary composite outcome



Primary endpoint composite of ischemic stroke/systemic embolism, major bleeding, or death



LAAO **Apixaban** **Warfarin**

Al-Abcha A, Saleh G, Sledge H, Abbasi M, Ismayl M, Killu AM, Simard T, Liu X, Hibbert B, Friedman PA, Siontis KC, Alkhouri M. Outcomes of stroke prevention strategies in patients with atrial fibrillation and end-stage renal disease. JACC Cardiovasc Interv 2025;18:2614-2624



CARDIAC ARRHYTHMIAS MANAGEMENT

- We suggest percutaneous left atrial appendage occlusion be considered in patients on PD with high risk of stroke but contraindications for long-term anticoagulant treatment ([2C](#)).

CONCLUSION

- What is new in the 2025 guidelines?



Key Changes

- Recommendation for maximizing loop diuretics with or without thiazide or thiazide-like diuretics to manage fluid overload.
- New recommendation regarding statin or statin/ezetimibe for secondary prevention of atherosclerotic cardiovascular disease.

Key Changes

- Revised, updated pharmacologic recommendations for heart failure management, including mineralocorticoid receptor antagonist and angiotensin receptor-neprilysin inhibitor.
- New recommendations regarding continuous glucose monitoring and glucose management.

Key Changes

- Revised, updated recommendation for direct oral anticoagulation for stroke prevention.
- New recommendations regarding percutaneous left atrial appendage occlusion for stroke prevention.

