Outcomes and care priorities for older people living with frailty and advanced chronic kidney disease: a multiprofessional scoping review

SUPPLEMENTARY DATA

Appendix 4: Table of Included Studies

	Aims	Design	Population	Frailty Assessments	PPI	Outcomes	Results
Baddour 2019 USA [1]	Determine the association of self-reported health and surprise question with frailty and functional status in older adults with chronic kidney disease (CKD) and their utility as screening tools	Prospective Cohort study	ACKD	Self-reported health, CFS, Fried, ADL's	N	Frailty and ADL'S	Both patient and provider subjective health assessments were moderately correlated with Fried frailty score, CFS score, and measures of functional status. SRH and the SQ demonstrated a fair correlation with each other. These findings were generally consistent regardless of age, the presence of diabetes or cardiovascular disease, or overall comorbidity burden.
Bancu 2017 Spain [2]	To define a frailty pattern among our dialysis population, to analyse the incidence and clinical evolution of these patients.	Retrospective cross sectional observational study	HD	Fried phenotype	N	Hospitalisation and mortality	Statistically significant differences between the two groups in terms of hospital admissions (0.77727) admissions/year of frail patients versus 0.2838 admissions per year of non-

Chang et al 2020	A randomized parallel	Protocol for	CKD3-5	Fried, hand grip	N		frail patients p = 0.005). Mortality in the group of frail patients was 20.45%, while in non-frail patients it was 12.36% (p < 0.005).
China [3]	controlled trial will be conducted to compare an individualized intervention according to the consequence of the comprehensive geriatric assessment with routine treatment.	RCT		strength, SPPB, MMSE			
Chao and Huang 2015 Taiwan [4]	We hypothesize that in ESRD patients, frailty might demonstrate significant association with their ECG parameters, thus potentially influencing the arrhythmogenic potential and the subsequent risk of cardiovascular mortality	Prospective Cohort study	HD	Six different self-report questionnaires with Chinese-translation including Strawbridge questionnaire (frailty if >1 positive domain), Edmonton frailty scale (if score ≥8), Simple FRAIL scale (if score ≥3), Groningen frailty instrument (if score ≥4), G8 questionnaire (if score 514), and Tilburg frailty instrument (if score ≥5)	N	ECG and QRS duration	Self-report frailty is significantly associated with QRS duration in chronic haemodialysis patients, and the association is independent of serum electrolyte levels and heart failure status

Chao et al 2015	Prospective cohort study	Prospective	HD	Strawbridge,	N	Frailty dialysis	Scores from each
Taiwan [5]	comparing self-reported	cohort study	110	Edmonton, G8,	'	complications	questionnaire showed
raman [5]	questionnaires	Conditional		Groningen, 5 item		Complications	significant association with
	demographic data			frailty Tilburg frailty			each other, except the G8
	collected and variables			indicator			questionnaire. FRAIL scale
	with complications of						correlated significantly with
	dialysis						age (P = 0.02), female gender
							(P = 0.03), higher Liu's
							comorbidity index (P = 0.02),
							lower serum albumin (P = 0.03)
							and creatinine levels (P < 0.01),
							and higher ferritin levels (P =
							0.02). Multivariate linear
							regression analysis identified
							an independently negative
							association between serum
							albumin and the simple FRAIL
							scale results (P = 0.01).
Chao and Huang	We hypothesize that GS	Prospective	HD	Frail scale	N	Medication	Using the simple FRAIL scale,
2016 Taiwan [6]	could modify the	Cohort study				adherence	10 (19.6%) patients were
	medication adherence					Taiwanese version	categorized as frail, and 24
	status of chronic dialysis					of the eight-item	(47.1%) were pre-frail.
	patients, and investigate					Morisky	Polypharmacy occurred in 48
	this issue using a					Medication	(94%) patients, and extreme
	prospectively enrolled					Adherence Scale	polypharmacy was found in 27
	cohort.					(MMAS)	(53%) patients. 80% of chronic
							dialysis patients had low
							medication adherence, and the
							severity of non-adherence
							increased significantly with
							younger age, lower degree of
							frailty, and lower dialysis
							clearance.
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Chao et al 2017 Taiwan [7]	Whether frailty is associated with temporal changes in BMD	Prospective Cohort study	ESKD	Frail scale	N	DEXA and bone health density	Among all ESRD survivors (mean 69.1 9.1 years, 47.2% male; frail 14%, pre-frail 53% robust 33%) The BMI among those with and without frailty no significant difference (p = 0:88).
Chao et al 2017 Taiwan [8]	We tested the hypothesis that self-report frailty increased the risk of incident Vascular Access (VA) failure after frailty assessment in patients.	Prospective Cohort study	ESKD	Self-reported frailty scale	N	Frailty, death, vascular access complications	Accounting for age, dialysis duration, DM, and laboratory results, Cox proportional regression analysis showed that frailty increased the risk of VA failure during follow up (hazard ratio [HR] 2.63, 95% confidence interval1.03–6.71, P = 0.04
Chao et al 2020 Taiwan [9]	A pilot study to validate the Laboratory deficit-based frailty index (LFI) against other frailty measures and examine outcome predictability (mortality)	Prospective Cohort study	ESKD	LFI and a modified Cardiovascular Health Study (CHS) scale based on self-report responses and six types of existing self- report frailty instruments in Chinese versions: the Strawbridge questionnaire (SQ), Edmonton frail scale (EFS), Groningen frailty indicator (GFI), Tilburg frailty indicator (TFI), G8 questionnaire and FRAIL scale	N	Biochemical and clinical variables	Frailty prevalence was 33.3% (CHS), 78.8% Strawbridge questionnaire, 45.5% (EFS), 57.6% (GFI), 27.3% (Tilburg frailty indicator), 84.8%(G8) and 18.2% (FRAIL) among ESRD participants. LFI-1 results were significantly correlated with those of LFI-2 (P< 0.01), EFS (P= 0.04) and GFI (P< 0.01), while LFI-2 results were not. Those with CHS or GFI-identified frailty had significantly lower 1,25-(OH)2-D levels than those without. After 5.4 months, patients with high LFI-1 scores, but not LFI-2, had higher mortality than those with lower scores.

Chowdhury et al 2017 Australia [10]	Articles before 2016 on frailty in CKD	Systematic Review	Pre dialysis, HD and KT		N	Minimental, ADL, iADL, Medicines	32 met the criteria. Twenty-three (72%) studies used or adapted the Fried phenotype to measure frailty. The prevalence of frailty ranged from 7% in community-dwellers (CKD Stages 1 –4) to 73% in a cohort of patients on haemodialysis. The incidence of frailty increased with reduced glomerular filtration rate. Frailty was associated with an increased risk of mortality and hospitalization.
Coelho 2020 Brazil [11]	AGNES is a prospective observational cohort that aims to investigate clinical, biochemical, and demographic factors associated with RRT initiation and mortality of patients with CKD stage 4 or 5 who are aged 70 years and older	Protocol	CKD 4-5	Full CGA, ADL's Cognition, Fried, CCI, Nutrition	N	Other measures sleep, monitoring device for BP and more biomarkers	N/A
Coppolino et al 2018 Italy [12]	To assess the entity of functional, general health and cognitive impairment and the possible relationship between these types of dysfunction and the severity of renal impairment.	Cross sectional	CKD	Fried phenotype, mini mental, ADL	N	Function, general health and cognition	Mild-to-moderate CKD is highly pervasive among frail elderly individuals and the severity of renal dysfunction is independently correlated with that of cognitive impairment.

Crespo et al 2020 Spain [13]	Incidence of COVID-19 in prevalent elderly KT patients, the characteristics of the first 16 symptomatic COVID-19 elderly KT recipients from a program and their outcomes until death or a minimum of 2 weeks after symptoms	Cohort	Transplant	Fried phenotype	N	Mortality	Short-term 14-day fatality rate in this group has been 50%, those who died had worse renal function before infection. They were more frequently obese, frail and had underlying heart disease.
De Sousa Meira et al 2016 Brazil [14]	Cross-sectional study to assess the frailty level of elderly under conservative management	Cross sectional	Conservative care	Edmonton Frailty Scale (0-17)	N	Socio- demographic and clinical variables	Minimum score of 1 and maximum of 14, with a mean score 7.71 (±3.10). Women (8.05±3.551) and illiterates (9.57±2.637) showed a higher mean score of frailty. By correlating the frail score, a moderate inverse correlation was found with years of study (p=0.033) and moderate positive correlation with some complications (p<0.05)
Drost et al 2016 Netherlands [15]	Cross-sectional study to measure the prevalence of frailty with two different measures	Cross sectional	ACKD,PD,HD	Frailty index (FI) and Frailty Phenotype (FP)	N	Prevalence of frailty	FI: 43.6% (>65) FP: No prevalence for over 65s provided. Being female and having more comorbidities was associated with higher frailty levels. The FI identified different but overlapping participants as frail compared with the FP; 62.5% of frail participants according to FI were also frail according to the FP

Farragher et al 2019 Canada [16]	The study objective was to formally assess the type and frequency of PD assistance received by patients over 50, and the relationship to observed frailty, functional status, and cognitive ability at the time of PD therapy initiation.	Prospective observational study	PD	Barthel, Lawton Brody IADL, MoCA, Fried, TUG, Hand grip strength	N	Peritoneal dialysis tasks were divided into 9 basic PD activities and 4 instrumental PD activities that need to be performed on a routine basis.	A total of 75 (62%) patients received assistance for a variety of tasks from friends or family (n = 41, 34%) or a paid caregiver (n = 34, 28%) 1 month after starting dialysis. At baseline, there was a high prevalence of functional dependency (79/120, 66%), frailty (71/110, 65%), and impaired cognition (68/115, 59%). Only 5% were fully independent, clinically robust, and scored within the normal range on cognitive testing. Factors associated with PD assistance included comorbidity (p < 0.03), cognitive impairment (p < 0.0001), and functional dependence (p < 0.02).
Fernandez et al 2019 US [17]	Multicentre prospective cohort study to test whether comorbidity is equally associated with waitlist mortality among frail and non-frail transplant candidates and to test whether measuring comorbidity burden and frailty improve mortality risk prediction.	Prospective cohort study	КТ	Fried phenotype	N	Transplant waitlist mortality	Among frail KT candidates, a high comorbidity burden is not associated with waitlist mortality. However, among candidates a high comorbidity burden is. These findings are consistent with the conceptual model for frailty in which this phenotype can result from comorbidity but also exist in the absence of comorbidity

Garcia-Canton et al 2019 Spain [18]	Observational prospective longitudinal study to estimate frailty prevalence in a haemodialysis population and its influence on short-term outcomes	Prospective observational study	HD	Edmonton Frailty Scale (EFS) (0-17)	N	Mortality, hospitalization and visits to hospital emergency services	According to the EFS, 82 patients (29.6%) were frail, 53 (19.1%) were vulnerable, and 142 (51.3%) were non-frail. During follow-up, 58.5% frail patients, 30.2% vulnerable, and 16.2% non-frail ones died (p<.005). A higher hazard of mortality was observed in frail than in non-frail patients (HR 2.34; 95% CI 1.39–3.95;p <.001). During follow-up the hospitalization rate was 852 episodes/1000 patient-years for frail patients, 784 episodes/1000 patient-years for vulnerable patients, and 417 episodes/1000 patient- years for non-frail patients
Gigilo et al 2015	Observational and cross-	Cross	HD	Fried phenotype	N	KDQOL, SGA and biochemical	(p<0005). The incidence ratio of visits to emergency services was 3216, 1735, and 1545 visits/1000 patient-years for each group (p<.001). Prevalence of frailty, prefrail
Brazil [19]	sectional study to examine the association between frailty and quality of life (QOL), nutritional status and clinical condition in elderly patients on haemodialysis (HD)	sectional				markers	and non-frail was 31% (n=48), 62% (n=97) and 8% (n=12), respectively. In conclusion elderly on HD frailty was associated with worse QOL and nutritional status. The inflammatory status was worse in the Frail and Pre-frail groups

Gopinathan et al	The aim of this study is to	Cross	HD	Fried SF 36, MoCA,	N	Frailty and	Prevalence of frailty was 56.4%
2020 India [20]	establish the prevalence of	sectional		self-reported frailty		cognitive	with method 1 and 64.1% with
	frailty in elderly patients					impairment	method 2. Patients having
	with CKD on HD,						cognitive impairment had a
	measured with two						significantly higher prevalence
	assessment instruments:						of frailty. The self-report
	physical performance						measure-based frailty
	measurement and self-						assessment (method 2) has a
	report measurement, in						lower cut-off point for the
	one population and to						definition of frailty as more
	determine the correlates						patients were identified as
	of frailty among the						frail, in comparison with the
	elderly patients on HD in						physical performance
	south India						measurement-based frailty
							assessment (method 1) in this
							dialysis population

Goto et al 2019	The aim of this study was	Cohort	HD	ADL's and iADL's,	N	ADL's and iADL's	Prevalence of functional
Netherlands [21]	to assess the association			Groningen Frailty		dependencies	dependence was high; four out
	at initiation of			Indicator GFI,		were categorized	of five participants were
	maintenance dialysis with			Caregivers received		in improvement	dependent in functional status
	functional status and			three questionnaires:		(score (> +1),	(30% ADL, 78% iADL) at
	caregiver burden.			the		stable (score of 0),	initiation of dialysis.
	Variables associated with			Neuropsychological		decline (score >-1),	Furthermore, almost half of
	functional change after			Inventory, the		and death.	the participants experienced
	initiating dialysis were also			Interview of			decline in functional status
	analysed. This was part of			Deterioration in Daily			(40%) or died (8%) within the
	larger prospective multi			Life Dementia , and			first 6 months after initiating
	centre cohort study			the Self-Perceived			dialysis. This decline was
	assessing the relationship			Pressure from			mostly due to loss of iADL
	of geriatric assessment			Informal Care (a			abilities (37% decline in IADL
	with outcomes in patients			Dutch questionnaire			versus 16% in ADL). Older age
	with ESKD			assessing caregiver			and a high score on the GFI
				burden).			were associated with the
							composite outcome functional
							decline/death. In addition, the
							percentage of caregivers
							reporting a high burden of
							care, increased from 23% to
							38%(P=0.004) after dialysis
							initiation

Goto et al 2019	The aim of this study was	Cohort	HD, PD,	7 domains were	Geriatric	Prevalence of geriatric
Netherlands [22]	to assess the prevalence		Conservative	assessed comorbidity	impairments	impairments was very high;
	of geriatric impairments		care	burden (Cumulative		77% of the patients had 2 or
	and frailty through a GA in			illness rating scale for		more geriatric impairments.
	a population with ESKD at			geriatrics, ADL,iADL,		Most frequent impairments
	the time of initiating			depressive symptoms		were seen in functional
	dialysis and in a			(GDS-15), nutrition		performance, cognition and
	population choosing			(Mini nutritional		severe comorbidity. Of the 89
	maximal conservative			assessment), mobility		conservative management
	management (MCM). Part			(TUG) and mini		population, the prevalence of
	of the larger GOLD study			mental HRQOL and		geriatric impairments was even
				other baseline		higher; 88% had 2 or more
				demographics were		geriatric impairments.
				done including frailty		
				screening using Fried		
				frailty index and		
				Groningen index		

Hall et al 2016	Two programs of care	Comparative	CKD	CGA-4 CKD Criteria	Geriatric	In both programs, at least 25%
USA [23]	were evaluated: A	study	CKD	and assessments of	assessments data,	of veterans had functional
U3A [25]		Study			,	
	geriatrician embedded in a			ADLs, iADLs, falls (self-	referrals, care	limitations that geriatric
	nephrology practice (CGA-			report of falls within	processes,	assessment identified.
	4-CKD Program) and a			prior 12 months),	qualitative themes	Cognitive impairment and
	nephrology clinic with			mobility, cognition		difficulty with iADLs were
	extended appointments			(dementia diagnosis		commonly identified. Geriatric
	for geriatric assessments			or Mini-Cog test),		assessments led to at least one
	(Renal Silver clinic).			frailty, Study of		care process in 45.4% (n = 15)
	,			Osteoporotic Fracture		of veterans in CGA-4-CKD and
				frailty criteria) and		37.1% (n = 13) of those in
				urinary incontinence.		Renal Silver. In Renal Silver,
				, , , , , , , , , , , , , , , , , , , ,		findings from 20.0% (n = 7) of
						geriatric assessments
						contributed to dialysis
						decision-making discussions
						that favoured conservative
						management (instead of
						dialysis). Additional care
						processes included social work
						referrals for assistance with
						advance directives or ADLs
						(17.1%, n = 6), and
						consultations for palliative care
						(8.6%, n = 3).
						• • • • • • • • • • • • • • • • • • •

Hall et al 2020 USA [24]	To identify quality of life themes that matter most to older adults receiving daily-sis and identify the extent to which existing quality of life instruments, specifically the KDQOL-36 and WHOQOL-OLD, overlap with those important themes.	Qualitative	HD	Screened for cognitive impairment and frailty using the mini-cog and a simple frailty questionnaire	Y	N/A	50% patients classed as frail. Two major quality of life themes were identified: (1) having physical well-being (2) having social support Perspectives on the subthemes often varied by frailty status. The majority of the subthemes did not correspond with domains in the KDQOL-36 and WHOQOL-OLD instruments.
Hernandez-Agudel et al 2021 Argentina [25]	To analyse the socio- demographic and clinical variables of patients who initiate haemodialysis or peritoneal dialysis due to end-stage kidney disease. The FRAIL scale was applied for the diagnosis of frailty syndrome	Cross sectional	HD + PD	CFS, hand grip, physical activity, social isolation	N	Socio demographic, biochemical and KT/V, hospitalisation	Prevalence of frailty in subjects who initiated renal replacement therapy was 55.55%, measured through the FRAIL Scale,higher in women than in men (p = 0.045). Frail patients had a higher Charleston comorbidity index (p =< 0.01). The mean serum creatinine, parathyroid hormone (PTH), and albumin were lower in frail patients, with statistically significant differences.
Hubbard and Peel 2016 Australia [26]	To determine whether the frailty status of patients with chronic kidney disease (CKD) can be measured using a Frailty index (FI)	Cross sectional	CKD	FI CKD, Fried	N	CGA	Mean FI-CKD was 0.25 (SD 0.12). The FI-CKD increased with age at 3% per year, correlated with a modified Fried phenotype (P < 0.001) and increased significantly across CKD stages (P = 0.04)

Hwang et al 2019 Korea [27]	Multicentre retrospective study comparing three risk stratification tools, to identify which is the best prognostic tool for predicting short-term mortality in elderly HD patients undergoing dialysis.	Retrospective	HD	CFS	N	Early mortality	The 3- and 6-month mortality rates were 31 (14.4%) and 48 (22.4%), respectively. Receiver operating characteristic curve analysis revealed that both score systems and the CFS showed similar performance while predicting 3- and 6-month mortality
lyasere et al 2016 UK [28]	To compare QoL of patients on assisted PD versus ICHD	Observational	Assisted PD and those on ICHD needing transport	CFS, minimental, falls and social	N	QoL assessments included Hospital Anxiety and Depression Scale (HADS), Short Form-12, Palliative Outcomes Symptom Scale (renal), Illness Intrusiveness Rating Scale, and Renal Treatment Satisfaction Questionnaire (RTSQ). Physical function was evaluated by Barthel Score and timed up and go test.	51.9% of aPD and 42.6% of HD patients were frail. Frailty was associated with worse SF-12 MCS, SF-12 PCS, Barthel Index, symptoms, illness intrusion, and HADS scores (P,0.01. In a posthoc analysis of 119 frail patients (frailty score 5) there was no significant difference in any QoL measure between assisted PD and HD after adjusting for the other covariates

lyasere et al 2019 UK [29]	To compare QoL across aAPD and HD	Observational	aAPD and HD	CFS and Barthel	N	HADS, SF-12 physical and mental scores, symptom score, Illness Intrusiveness Rating Scale (IIRS), Barthel's score, and the Renal Treatment Satisfaction Questionnaire (RTSQ)	There was no statistically significant difference in any of the outcome measures between HD and PD. Longitudinal trends in outcomes were also not significantly different.
lyasere et al 2019 UK [30]	To compare QoL of conservative care management patients with aPD and ICHD patients	Observational cohort	Conservative management ICHD and aPD	CFS	N	Short Form-12 [SF12; Physical Component Summary scale (PCS) and Mental Component Summary scale (MCS)], HADS, Palliative Outcomes Scale- Symptoms (POS-S) (Renal), Illness Intrusiveness Rating Scale (IIRS), Renal Treatment Satisfaction Questionnaire (RTSQ) and the Barthel score	Frailty CCM: 39.3%; aPD: 60.7% ICHD: 39.3%. Frailty was associated with lower PCS on SF12, higher depression score and illness intrusiveness.

Johansen et al 2007 USA [31]	To explore the prevalence, predictors and outcomes of frailty	Cohort	HD	Questionnaire based frailty from Women's Health Initiative cohort (see text). Measuring physical functioning, fatigue, activity, nutrition	N	Mortality and hospitalisation	75% of patients over 60 are frail. Older age, female sex, and haemodialysis (rather than peritoneal dialysis) were independently associated with frailty. Coxproportional hazards modelling indicated that frailty was independently associated with higher risk of death and with the combined outcome of death or hospitalization
Kakio et al 2018 Japan [32]	Examine association between diabetic nephropathy and frailty	Cross sectional observational	HD	Fried	N	Frailty and diabetic nephropathy	21.3% were frail, 51.6% were prefrail. The prevalence of frailty in the Diabetic Nephropathy group was significantly higher.
Kallenberg et al 2016 [33]	Articles until 2016, reporting on association of functional or cognitive impairment or frailty with adverse outcome	Systematic Review			N	Geriatric impairments and outcomes	30 articles were identified that reported on 35 associations. Mean age was >60years old in 73% of the studies, and geriatric conditions were highly prevalent. Twenty-four studies (80%) reported on functional impairment, seven (23%) reported on cognitive impairment, and four (13%) reported on frailty. Mortality was the main outcome measure in 29 studies (97%), and one study assessed functional status trajectory. In 34 of 35 (97%) associations reported, functional or cognitive impairment or frailty was significantly and

							independently associated with adverse health outcomes.
Kamijo et al 2018 Japan [34]	To evaluate whether sarcopenia and frailty have a negative impact on mortality in PD patients. Also to investigate the morbidity of sarcopenia and frailty, and assessed the factors related to disease conditions, such as malnutrition-inflammation-atherosclerosis (MIA) syndrome	Prospective Cohort study	PD	CFS	N	Mortality	13.1% of patients >60 was classed as frail. Patients with frailty are more likely to be older and exhibit higher BI and CCI values, as well as lower PS, SMI, handgrip strength, and usual walking speed values. After adjustments were made for potential con-founders, including age, CFS, gender, walking speed, SMI, and grip strength, the CFS remained an independent predictor of mortality.
Kuki et al 2020 Japan [35]	This study aimed to prospectively evaluate whether the two physical performance parameters are associated with the onset of fatal/non-fatal CV events in patients undergoing HD	Prospective cohort	HD	Gait speed and HGS	N	CV events	Among Japanese HD outpatients who were able to walk independently, slow usual GS and weak HGS were significantly associated with CV events, independent of age, sex, HD duration, and medical history
Kumarasinghe et al 2021 Australia [36]	The aim of the study was first to assess the feasibility of incorporating CFS assessment into routine outpatient nephrology practice in the pre-dialysis setting and then to explore the association of the degree of frailty with planned ESKD management.	QI project	Pre-dialysis	CFS	N	The primary outcome assessed was feasibility of CFS assessment. The secondary outcomes were associations of frailty assessment with planned ESKD management and clinician perception of the	21% of patients who had completed CFS' were planned for conservative management in contrast to only 5% of those who had 'no CFS completed' (P = 0.013). Even when adjusted for age, gender, assessing clinician, usual nephrologist and clinic setting, the 'completed CFS' group was more likely to be planned for conservative management (P =

						utility of frailty assessment.	0.026). The 'completed CFS' group was less likely to have haemodialysis (23%) compared with those with 'no CFS' (40%) (P = 0.037). Completed CFS group was more likely to have ESKD management plan 'not documented' (21%) than the 'no CFS' group.
Kutner et al 2014 USA [37]	Aim was to review the Fried Frailty prevalence in large cohort of dialysis patients- this was part of larger study titled ACIVE-ADIPOSE part of USRDS.	Cohort	HD	Fried	N	CV and other parameters	In the > 65 years 45% of this group identified as frail. Increased association with frailty and CVD and PVD
Lee S.J et al 2015 South Korea [38]	The purpose of this study was to compare the factors that influence frailty and to investigate the contribution of frailty to the HRQOL of predialysis CKD patients in Korea. A cross sectional survey	Cross sectional survey	CKD 2-4	Frailty was identified using physical functioning and weakness, then SF36 used as questionnaire for HRQOL	N	HRQOL	37.5% were frail, all older. Frail patients had lower physical and mental quality of life. However, gender, educational level, marital status, CKD stage, average income, blood pressure, BUN, creatinine, and haemoglobin level were not associated with frailty. Influence of frailty on health-related quality of life

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Lee S.W et al 2017	To identify the association	Prospective	HD	For frailty assessment	N	The outcome was	In multivariate logistic
South Korea [39]	between frailty and	observational		CGA protocol used.		the composite of	regression analysis, after
	adverse outcomes, and	cohort		Using the scores of		all-cause death or	adjusting for age, sex, diabetes,
	the change of frailty			nine items in CGA		cardiovascular	body mass index (BMI), and
	before and after dialysis			protocol: diagnosis of		hospitalization.	time of predialytic nephrologic
	initiation.			malignancy,			care, female sex, and increased
				Charleston			BMI were associated with
				comorbidity index,			increased and decreased odds
				serum albumin levels			of frailty, respectively. In
				and IADL, Korean			multivariate Cox proportional
				Mini-Mental State			hazards analysis, after
				Examination score,			adjusting for age, sex, diabetes,
				Nursing Delirium			BMI, and time of pre-dialytic
				Screening scale,			nephrologic care, frailty was
				malnutrition based on			significantly associated with
				the Mini Nutritional			the composite adverse
				Assessment (MNA)			outcome. In repeated frailty
				score, and mid-arm			assessments, the
				circumference.			multidimensional frailty score
							significantly improved 12
							months after the initiation of
							dialysis, which largely relied on
							improved nutrition.
Lin Hsin-Chang et	The goal of this study is to	Retrospective	HD	5 Point frailty	N	Vascular access	Age, fatigue, and weight loss
al 2020 Taiwan	evaluate the factors,	review		,		failure, KT/V	are useful prognostic indicators
[40]	including frailty status and						for the identification of
[]	vascular access						recurrent VAF.
	characteristics,						
	demographic data, BMI,						
	arteriovenous fistula						
	functions, and						
	biochemistry, that may be						
	predictors for recurrent						
	episodes among elderly						
	dialysis patients who have						
	experienced previous VAF.						
	experienced previous VAF.						

Lo et al 2008	A Prospective Pilot Study	Prospective	HD admitted	Version of Fried	N	Tests used	A total of 73.3% of patients
Canada [41]	to Measure Changes in	Pilot	acutely	version of Fried	IN .	included the 4-	(95% confidence interval, 54.1
Callaua [41]	Functional Status	FIIOL	acutely			item Basic Activity	to 87.7) experienced a decline
	Associated With					•	in personal functional
						of Daily Living	1 .
	Hospitalization in Elderly					(BADL) measure	independence in association
	Dialysis-Dependent					and the Lawton-	with hospitalization.
	Patients. To determine					Brody Scale of	
	functional impairment at					Instrumental	
	the time of admission and					Activities of Daily	
	again at 1 week after					Living (IADL).	
	discharge					Physical	
						performance was	
						measured using	
						the timed up-and-	
						go (TUG) test and	
						grip strength. In	
						addition, cognitive	
						function was	
						measured using	
						the Trails A & B	
						tests and the clock	
						test.	
Lopez-Montes et	Study to analyse	Cohort	HD	Fried	N	Mortality;	53.8% of patients were frail.
al 2020 Spain [42]	depression, cognition, and					disability in basic	Frail older adults that initiate
	physical function change					and iADL, physical	haemodialysis present higher
	in older adults on					function with	mortality than the non-frail
	haemodialysis at 12-					SPPB, Mini	ones at 12-month follow-up.
	month follow-up,					Cognitive	Frail patients that survive
	depending on frailty					Examination,	improve physical function,
	status.					Geriatric	depression and inflammatory
	status.					Depression Scale	profile compared to the non
						(GD).	frail ones
						• •	i ali olles
						Inflammatory and	
						nutrition profile	

Mc Adams-	Whether frailty measured	Prospective	KT	Fried	N	LOS post	Frailty was independently
DeMarco 2017	pre kidney transplant (KT)	Cohort	KI	THEU	'\	transplant	associated with longer LOS
USA [43]	was associated with LOS	Conort				transplant	(RR=1.15, 95%CI: 1.03-1.29;
U3A [43]	was associated with EO3						P=0.01) and LOS≥2 weeks
							(OR=1.57, 95%CI:1.06-2.33;
							P=0.03) after accounting for
							registry-based risk factors,
							including DGF. Frailty also
							,
							attenuated the association
							between LOS and mortality (
							HR:1.55 95%CI:1.30-1.86,
							P<0.001; frail HR=0.97,
							95%CI:0.79-1.19, P=0.80; P for
NA - A - I	4)	Dun ou outing	L/T	Falled	N.	ADI /IADI	interaction=0.001).
McAdams-	1)Identify characteristics	Prospective	KT	Fried	N	ADL/IADL	Frailty was 23.7% among those
DeMarco et al	of frail KT recipients, 2)	cohort				disability, CESD	aged 65-74 years, and 22.7%
2017 USA [44]	identify the most common					depression,	for those recipients aged ≥75
	components of frailty					education, and	years. Pre-frailty was 38.1 for
	among KT recipients and					HRQOL	>65s and 50% for >75s. Older
	3) explore which patterns						Age, IADL disability, depressive
	of the frailty components						symptoms, less than a high
	are most strongly						school education and low
	associated with mortality						HRQOL predicted frailty. The
	risk among KT recipients						most common frailty pattern
							was poor grip strength, low
							physical activity and slowed
							walk speed. KT recipients with
							exhaustion and slowed walking
							speed and poor grip strength,
							exhaustion, and slowed
							walking speed were at
							increased mortality risk.

Mei et al 2021	The inclusion criteria were	Systematic			N/A		Frailty is a risk factor for
China [45]	as follows: (1)	Review and			,		mortality, hospitalization, and
	retrospective and	meta-analysis					falls, indicating it is important
	prospective cohort	,					to assess frailty as a prognostic
	studies; (2) patients with						factor for CKD. Included were
	clinically confirmed CKD;						recent cohort studies using a
	(3) frailty was defined by a						broader definition of frailty
	recognized criterion and						and numerous negative health
	using the established						outcomes, because
	frailty models or modified						assessments of frailty are
	versions; (4) outcomes						generally correlated. Subgroup
	were all-cause mortality,						analysis based on possible
	all-cause hospitalizations,						influencing factors to
	and falls						comprehensively understand
							the effect of frailty on risk
							factors and prognosis, which
							may provide clinicians with an
							indicator to measure the
							severity, prognosis, and
							progression or reversal of CKD
Meulendijks et al	Aim to assess whether the	Prospective	Pre-dialysis	GFI is a 15 item frailty	N	Referral to	Geriatric impairments were
2015 Netherlands	Groningen frailty indicator	Cohort		assessment (4		geriatrics,	prevalent with percentages
[46]	(GFI) can be used to			geriatric domains-)		mortality,	ranging from 23% (visual
	distinguish fit older ESRD			done at time of		hospitalisation	impairment) to 62% (hearing
	patients, likely able to			referral into pre-			impairment). One-third of
	tolerate and benefit from			dialysis clinic. If			patients were identified as frail
	dialysis, from frail older			scored high referred			according to the GFI, (GFI_4)
	patients who need further			to geriatric nurse and			and these patients more
	evaluation with a			geriatrician			frequently required
	geriatrician's						hospitalization. Both the GFI
	comprehensive						and the nephrologists'
	assessment						assessment failed to identify
							relevant geriatric impairments
							that were detected after the
							geriatric consultation.

Moffatt et al 2018	Qualitative study to 1)	Qualitative	Nurses	Frailty Assessment for	N	N/A	Nurse participants reported an
Canada [47]	explore the nurse			Care Planning Tool/		,	overall positive experience
	experience of screening			Clinical Frailty Scale			using the FACT method to
	for frailty using the frailty			,			screen for frailty and indicated
	assessment for care						that their understanding of the
	planning tool (FACT) tool						multiple dimensions and
	2) determine how, if at all,						subtleties of "frailty" were
	provider perceptions of						enhanced. Future nurse-led
	frailty changed after						FACT screening initiatives
	implementation of the						should incorporate those
	frailty screening tool; and						factors identified as being
	3) determine the						integral to program success:
	perceived factors that						realistic goals, clear guidelines,
	influence uptake and						and ongoing training.
	administration of the FACT						
	screening tool.						
Moreno-Useche	The aim of this research is	Cross	HD/PD	Self-reported Frail		Biochemical, social	Prevalence of frailty syndrome
et al 2020	to analyse the socio-	sectional		scale		and frailty	in subjects who initiated renal
Columbia [48]	demographic and clinical						replacement therapy was
	variables of patients who						55.55%, measured through the
	initiate haemodialysis or						FRAIL Scale, The prevalence of
	peritoneal dialysis due to						frailty syndrome was higher in
	end-stage kidney disease.						women than in men (p =
	The FRAIL scale was						0.045). Frail patients had a
	applied for the diagnosis						higher Charleston comorbidity
	of frailty syndrome						index (p =< 0.01). The mean
							serum creatinine, parathyroid
							hormone (PTH), and albumin
							were lower in frail patients,
							with statistically significant
							differences.

Malanata at al	O	C	LID	Esta d	N.	Dia ad la a a ad	The DC1
Nakazota et al	Our aims for this study	Cross	HD	Fried	N	Blood based	The PC1 score was associated
2020 Japan [49]	were (1) to examine	sectional				parameters	with the prevalence of frailty
	whether the PC1 score is						and was an independent
	an appropriate measure of						predictor for frailty (odds ratio
	physiological						per SD: 2.31, P=0.01) using a
	dysregulation and frailty,						multivariate logistic regression
	and (2) to validate our						model, which showed good
	interpretation of						discrimination (c-statistic:
	biomarker variability.						0.85). Therefore, the PC1 score
							represents principal
							information shared by
							biomarker variabilities and is a
							reasonable measure of
							homeostatic dysregulation and
							frailty
Neradova et al	The aim of this study was	Retrospective	Dialysis,	CFS	N	Mortality and	In the 174 patients who had a
2021 UK/Brazil	to determine whether pre-	cohort study	CKD, KT			hospitalisation	CFS recorded, the age was 65.4
[50]	existing frailty was a risk	,					years ± 15.8 (mean ± SD) and
	factor for hospital						57,5% were male. At the end of
	admissions and mortality						follow up, 26% had died. Frail
	in renal patients in the						patients (CFS 5-7) were more
	first wave of the COVID						than three times more likely to
	pandemic.						die compared to less frail
							patients (CFS of 1-4) (odds
							ratio (OR) 3.3, 95% confidence
							interval (CI) 1.0-10.6). 118
							patients (68%) required
							admission, but there was no
							difference in hospital
							admission rates for frail vs non-
							frail patients (OR 0.6, CI 0.3-
							1.7).

Nixon et al 2019 UK [51]	An evaluation of the diagnostic accuracy of several frailty screening methods	Convenience series	CKD 4-5	Baseline data , Charleston comorbidity, karnofsky performance status, minimental risk evaluation for eating and nutrition. Frailty screening methods CFS, PRISMA 7, CKD FI, CKD FI LAB, walking speed, hand grip strength and short physical performance battery (SPPB). Frailty phenotype used as standard reference	N	Clinical Frailty Scale, PRISMA-7, CKD Frailty Index, CKD FILAB, walking speed, hand grip strength and Short Physical Performance Battery	Overall walking speed had the highest accuracy for physical screening, The CFS had the highest AUC for non physical screening in this population. Nineteen (21%) patients were categorised as frail, 42 (47%) as pre-frail and 29 (32%) as robust.
Nixon et al 2020 UK [52]	The EX-FRAIL CKD trial aims to inform the design of a definitive randomised controlled trial (RCT) that investigates the effectiveness of a progressive, multicomponent homebased exercise programme in prefrail and frail older adults with CKD.	Protocol 2 arm parallel RCT	CKD 3b-5	CFS	N	Primary feasibility outcome measures include rate of recruitment, intervention adherence, outcome measure completion and participant attrition.	N/A
Nixon et al 2020 UK [53]	This study aimed to evaluate the relationship between frailty and HRQOL in patients with CKD Stages 4 and 5 (G4–5)	Secondary analysis	CKD4-5/HD	Fried	N	QOL	Frailty is independently associated with worse HRQOL in patients with CKD G4–5D. Exhaustion, or fatigue, is the most significant Frailty Phenotype component

	and those established on haemodialysis (G5D).						contributing to worse HRQOL in those with advanced CKD
Nixon et al 2021 UK [54]	The aims of this quality improvement project were to: (1) proactively identify people living with frailty and CKD; (2) introduce a practical assessment, using the principles of the comprehensive geriatric assessment (CGA), for people living with frailty and chronic kidney disease (CKD) able to identify problems; and (3) introduce person-centred management plans for people living with frailty and CKD.	QI project	CKD/HD	CFS/GA	N	Hospitalisation and mortality, number and type of problems encountered	One hundred and fifty patients (33%) were screened as frail (i.e. CFS ≥ 5). Frail patients were older than non-frail patients (median age 81 years [IQR 14] vs. 74 years [IQR 11], p < 0.001). This quality improvement project demonstrates the high burden of frailty and problems experienced by those with CKD. It also details an approach to the implementation of a frailty screening programme and GA service so that problems can be identified and a person-centred management plan developed.

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Nixon et al 2021	The aim of the EX-FRAIL	Pilot RCT	CKD 3b-5	SPBB, FP, Barthel	N	Primary outcome	A median of 28 (IQR 16)
UK [55]	CKD Trial, a pilot			index, FES-1, RENAL		measures included	exercise sessions were
	randomised controlled			POS, Short SF -12		eligibility,	completed during the 12-week
	trial (RCT), is to inform the					recruitment,	intervention period. Eleven
	design of a definitive RCT					intervention	(73%; 95% CI 45, 91) exercise
	that evaluates the					adherence,	group participants completed 2
	effectiveness of a home-					outcome measure	exercise sessions per week,
	based exercise					completion and	with a mean of 36.5±8.5
	intervention in pre-frail					participant	minutes spent exercising each
	and frail older adults with					attrition rate.	session. The mean RPE score
	CKD					Secondary	for exercises 2–6 was 12±2.
						outcomes PF,	The main reasons for missing
						Frailty, ADL. Falls,	exercise sessions were pain (n
						Symptom burden,	= 14; 56%; 95% CI 35, 75),
						HRQOL	participant wishes (n = 7; 28%;
						1111402	95% CI 13, 50) and feeling
							unwell (n = 4; 16%; 95% CI 5,
							37).
							37).

Novais et al 2020	The aim of this study was	Observational	CKD and	Cumulative illness	N	Geriatrician	On univariate analysis, the
France [56]	to assess the prevalence	retrospective	dialysis	rating scale, MNA,		recommendations	geriatric impairments
	of geriatric impairment	study	,	cognition SPPB, Fried,		for KT	influencing the geriatricians'
	and frailty in older	,		polypharmacy			recommendations for KT were
	patients with advanced						multiple comorbidities,
	CKD using a pretransplant						cognitive and physical
	CGA model. The secondary						impairment, symptoms of
	objectives of this study						depression, nutritional status,
	were to compare the						and dependence on ADL and
	geriatric impairments						IADL
	between dialysis and non						
	dialysis patients and to						
	identify the main geriatric						
	impairments influencing						
	the geriatricians'						
	recommendations for KT.						
Orlandi &	To assess frailty levels of	Cross	HD	Edmonton Frailty	N	Frailty and falls	Around 35.0% of the elderly
Gesualdo et al	elderly HD patients	sectional		Scale (0-17)			showed no frailty, 26.7% were
2014 Brazil [57]		study					vulnerable, 20.0% showed mild
							frailty, 13.3% moderate frailty
							and 5.0% severe frailty.
Painter et al 2013	To compare two methods	Cross	HD	Fried , Physical	N	Sub analysis of	The proportion of patients in
USA [58]	to operationalise fried	sectional		function in SF-36		renal exercise	the REXDP who met the
	phenotype frailty scores 1.					demonstration	criterion of frailty was 24%
	Using actual measures of					study	when actual measures were
	walking speed and						used for walking speed and
	strength (FRAIL meas). 2.						weakness components. It was
	The substitution of the PF						higher in the older > 65 year
	scale for walking speed						age group The correlation
	and strength (FRAILsubst).						between gait speed and the PF
	A second aim of this						score was 0.34 and the grip
	analysis was to identify						strength measures were 0.14,
	clinical and self-reported						both of which were statistically
	health-related quality of						significant
	life (HRQoL) factors that						

	are associated with frailty in haemodialysis patients.						
Parlevliet et al 2012 Netherlands [59]	Cross sectional multicentre study to assess the use of a CGA on randomly selected ESRD patients for prevalence and feasibility. Secondary aims were to compare to similar cohort of cancer patients	Cross sectional	HD and PD	A full CGA including all four domains (plus A DL's) completed by face to face interview in the patients home, followed by interviews with the MDT on acceptability and relevance of CGA questionnaire	Υ	Domains of CGA, feasibility of implementation as perceived by professionals, patients and carers	The most prevalent geriatric condition was polypharmacy (94.6%). Frequently observed were hearing impairment (36.8%), malnourishment (32.7%), social or emotional loneliness (30.6% combined) and depressive symptoms (24.5%). 24.0% o reported pain. Reported difficulties related to housekeeping, travelling and walking. Caregivers reported a number of behavioural problems eating behaviour most prevalent (34.0%). 84.4% of caregivers experienced care as a large burden.

Perez-Saez et al 2021 Spain [60]	To study the potential effects of multimodal rehabilitation as a prognostic variable to predict the 90-day primary endpoint based on clinical and functional outcomes achieved in frail and nonfrail KT candidates	Protocol RCT	KT	Fried phenotype, frail scale, CFS, Edmonton frail scale, Frail-VIG index		The primary endpoint will be a composite achievement of clinical and functional main outcomes in frail and non-frail KT candidates at 90 days post-transplantation.	N/A
Pugh et al 2016 UK [61]	The aim of the current study was to assess the impact of both frailty and comorbidity on the outcome of patients referred to the pre-dialysis service.	Observational	Pre dialysis.	CFS CCI	N	Mortality	Statistically significant difference between the groups, with the increasingly frail or comorbid subgroups having the worst survival. The percentage of patients dying within 3 years of the home visit increased with each increase in CFS score. Patients with higher levels of frailty were more likely to choose conservative care

Pyrat et al 2020 UK [62]	We sought to describe the choices and outcomes of a large cohort of older predialysis patients who underwent pre-dialysis education in our centre	Cohort	CKD-5 Pre dialysis	CFS	N	Patient characteristics influenced whether patients chose to receive HD,PD, HHD, transplant or opted for MCM. Impact of patient characteristics and treatment decisions on survival were compared from the time of final choice	By adding CFS to the model (n = 486)—sex, age, choice and CFS all predicted 3-year survival while comorbidity had no significant effect. However, when only patients with CFS 6 (at least moderate frailty, needing assistance with all outside activities or unable to undertake outside activities without support) were included in the analysis (n = 140, MCM 83 and HD 57), only age rather than the choice of RRT vs MCM significantly affected survival
Reece et al 2013 USA [63]	1) determine if greater CKD severity was independently associated with the end points of poor physical performance and frailty in a large, racially diverse sample of adults with CKD, and (2) identify additional risk factors for poor physical performance and frailty in CKD.	Cross sectional	CKD 4-5	Fried	N	Fried frailty measure, SPPB	Worse renal function demonstrated a graded association with worse physical performance and frailty, independent of age, sex, race, BMI, anaemia, socioeconomic status and major comorbidities

Salter et al 2015 USA [64]	A cross sectional study examining perceived frailty, patient nurse and nephrologists.	Cross sectional	HD	Fried criteria. Perceived frailty documented by patient, nephrologist and nurse, patients also asked 'how frail do you think you are'	N	Perceived and measured frailty	In multivariable models, only disability was associated with measured frailty (adjusted OR [aOR] = 1.47, 95% CI: 1.04-2.08, P = 0.03 for each additional ADL difficulty). Among frail participants, only 42.0% and 39.2% were correctly perceived as frail by their nephrologist or NP, and only 4.9% perceived themselves as frail Older adults were more likely to be
Sclauzero et al 2013 Italy [65]	The study aimed to evaluate the effect of the different components of frailty on the QoL of people on dialysis.	Cross sectional	HD	ADL's, Cognition, Karnofsky Index, SF 36, SGA	N	QOL	perceived as frail by nephrologist but less likely by themselves. 32.5% of patients revealed one or more disabilities (ADL scale) and that 38.4% were totally or partially dependent (IADL scale). KI demonstrated that 42.9% of subjects needs help to take care of themselves. Malnutrition was present in 34% of the subjects investigated.
Song et al 2020 China [66]	Inclusion criteria: (1) risk factors for mortality of elderly haemodialysis patients were the subject; (2) haemo dialysis patients included an elderly population; and (3) study data included odds ratio(OR) values and 95% confidence intervals (CIs)	Systematic Review and meta-analysis	HD		N	Geriatric impairments	Functional impairment (OR = 1.45, 95% CI: 1.20–1.75), cognitive impairment (OR = 1.46, 95% CI: 1.32–1.62) and falls (OR = 1.14, 95% CI: 1.06–1.23) were significantly and independently associated with increased mortality in elderly haemodialysis patients

	or data that could be transformed to OR values and 95% CIs by statistical methods.						
Suzuki et al 2019 Japan [67]	Effects of electrical muscle stimulation in frail elderly patients during haemodialysis (DIAL): rationale and protocol for a crossover randomised controlled trial	Protocol Intervention RCT	HD	SPBB, Mini Cog, Nutrition	N	Change in quadriceps isometric strength after intervention, improvement in physical function.	N/A
Takeuchi et al 2018 Japan [68]	Cross-sectional study to determine the prevalence of frailty and associated factors	Cross sectional	HD	Fried and self- reported frailty	N		Frail (over 65s): 26.1% over 75s: 32.7%. Pre-Frail over 65s: 51%, over 75s: 56.6%. Frailty was significantly associated with female gender, age, age >75, BMI<18.5, number of medications, diabetes and nutrition status (MNA-SF).
Tan et al 2018 Australia [69]	Prospective study to explore the impact of initiating dialysis on symptom burden and functional status as well as its trajectory as dialysis progresses in the elderly patient.	Prospective cohort	Supportive care, HD, PD	Fried	N	Symptom burden	The commencement of dialysis in a younger cohort of elderly patients was associated with a small but significant decrease in overall symptom burden. There was no change in functional status in both groups over a six-month follow-up period.

Van Loon et al	The aim of this review is to	Systematic		N	N/A	Only 7 studies carried out an
2016 Netherlands	give an overview of all	review				analysis of elderly patients
[70]	currently available					(>70 years old). Malnutrition
	evidence regarding the					and frailty were systematically
	relation of geriatric					assessed, and their relation
	impairments and the					with mortality was clear
	accumulation of					although it is lacking in the
	impairment across these					elderly population specifically.
	domains at initiation of					In addition, cognitive
	dialysis with mortality and					impairment and functional
	dialysis-related					outcomes at the initiation of
	complications.					dialysis were related to an
						increased mortality in most
						studies. However, not all
						studies applied systematic
						assessment tools, thereby
						potentially missing relevant
						impairment. None of the
						studies applied a geriatric
						assessment across multiple
						domains.

Van Loon et al 2017 Netherlands [71]	Cross sectional study of incident dialysis patients part of the GOLD study. Consecutive patients eligible for dialysis were included between 3 weeks before and 2 weeks after the first dialysis session.	Cross sectional	HD	ADL's, iADL's, Timed-Up-and-Go, the Geriatric Depression Scale, nutrition (the MiniNutritional Assessment, comorbidity burden. The cognitive test battery. In addition: the Groningen Frailty Indicator, the Fried Frailty Index, Geriatric8, the Identification of Seniors at Risk-Hospitalized Patients screening, the Hospital Safety Program criteria and the clinical judgment of the nephrologist (frailty question).	N	Geriatric assessment, frailty and clinician view	Functional impairment was high: 78% needed help with iADL's, 34% needed help with basic ADL, and in 25% of patients, mobility was severely impaired. Cognitive impairment was present in 66% of patients. Severe comorbidity burden in 35% of patients. Of all 75% (n=92) had impairment in two or more domains and were considered frail. lowest percentage of frailty was found with the Fried Frailty Index (48%), and the highest percentage was with Geriatric8 (88%). Not one showed overall 100% sensitivity. Of the frailty screening tools, the Identification of Seniors at Risk has the best discriminating abilities in the ESRD population, showing the highest specificity and a fairly good sensitivity, and 91% of
				I =			has the best discriminating abilities in the ESRD population, showing the highest specificity and a fairly

Van Loon et al	To assess quality of life in	Cohort	Pre dialysis	Frailty as measured by	N	QOL	Frailty in 88% of patients who
2019 Netherlands	patients starting dialysis	Conort	Fie dialysis	geriatric screening	IN	QOL	chose conservative care, 78%
	1.			genatric screening			I
[72]	and patients choosing						who chose dialysis. Baseline
	maximal conservative						QoL did not differ significantly
	care.						between the groups. After six
							months, EQ-5D Index did not
							improve significantly in the
							dialysis group, but a small but
							clinically relevant decline was
							seen in the conservative group.
							Hospitalization occurred in
							50% of dialysis patients vs. 24%
							of conservative patients (p<
							0.01). In the extended 12-
							months analysis, mortality rate
							in conservative patients was
							34% compared to 16% in
							dialysis patients (p= 0.01). In
							patients over 80 years old, no
							survival benefit could be found
							for dialysis patients starting
							dialysis vs. MCM.
Van Loon et al	Multicentre, prospective,	Prospective	Pre-dialysis	Geriatric assessment:	N	Mortality and	Twelve-month mortality risk
2019 Netherlands	cohort study assessing the	cohort		including assessment	'	hospitalisation	was higher in patients with ≥3
[73]	relation between a GA and			of (instrumental)			impairments (hazard ratio [HR]
[,3]	poor outcome in ESKD			activities of daily			2.97 [95% CI 1.19–7.45])
	patients.			living (ADL), mobility,			compared to less impaired
	patients.			cognition, mood,			patients. FFI frail patients had a
				nutrition, and			higher risk of 12-month
				comorbidity. In			mortality (HR 7.22 [95% CI
				addition, a frailty			2.47–21.13]) and
				screening (Fried			hospitalization (OR 1.93 [95%
				<u> </u>			, , , , , , , , , , , , , , , , , , , ,
				Frailty Index, [FFI])			CI 1.00–3.72]) compared to fit
		1		was applied			patients.

Van Loon et al	To assess the prevalence	Longitudinal	HD,PD aPD	Canadian Study of	N	Falls, other	Mean frailty score 4.3 (SD1.2),
2019 UK [74]	of falls and the impact on	observational	TID,FD aFD	Health and Aging	l IN	outcomes were	frailty was slightly higher in
2019 OK [74]	mortality and quality of	study		Scale		ALD, HADS, QOL	aPD group. patients with
	life in frail elderly patients	Study		Scale		SF-12	diabetes mellitus were twice as
	on assisted PD (aPD) and					3F-1Z	likely to report falls at baseline
	haemodialysis (HD) from						and falls at baseline were
	the FEPOD Study.						associated with falls during
	the FEFOD Study.						follow-up. Literature revealed
							frailty was a strong risk factor
							for falling and falling results in
							a higher mortality and
							hospitalization rate. Fall
							incidence was comparable in aPD and HD.
Van Munster et al	Prospective cohort study	Draspastiva	HD,PD Pre	FI, GFI self reported	N		Different short questionnaire—
2016 Netherlands	of prevalence of frailty by	Prospective cohort		· •	l IN		•
		COHOIT	dialysis	items, the VMS (Dutch assessment),			based screening instruments, i.e. the GFI, ISAR-HP and VMS,
[75]	screening using different instruments			ISAR-HP			1
	instruments			ISAK-TP			can all be applied to the ESRD population to screen for frailty
							in both young and older ESRD
							patients. Of these three
							1 .
							screening instruments, the GFI
							showed the highest sensitivity
							and negative predictive value
							for screening frailty in dialysis
							and pre-dialysis patients, with
							the FI as the gold standard. In
							addition, the ISAR-HP also
							showed comparable
							performance to that of the GFI,
							with slightly lower negative
							predictive value, and had the
							highest positive predictive
							value of all three instruments.

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Van Oevelen et al	A prospective,	Protocol	CKD 5	Nephrogeriatric	N	The primary	N/A
2021 Netherlands	observational cohort	prospective		assessments		outcome is HRQoL,	
[76]	study. Initial stage when	cohort				measured with the	
	dialysis is initiated or eGFR					Twelve-item Short-	
	drops to 10					Form Health	
	mL/min/1.73m2 or lower,					Survey. Secondary	
	the second stage of the					outcomes are	
	study commences. In both					clinical outcomes	
	stages nephrogeriatric					(mortality,	
	assessments will be					hospitalisation,	
	performed annually,					functional status,	
	consisting of					cognitive	
	questionnaires and tests					functioning,	
	to assess most common					frailty), cost	
	geriatric domains, i.e.					effectiveness, and	
	functional, psychological,					decisional regret	
	somatic, and social status.						
Vettoretti et al	We evaluated if Frailty	Cross	Pre dialysis	Fried , nutritional	N	Fried phenotype	Frail patients (F-CKD) had
2020 Italy [77]	Phenotype (FP) could	sectional	-	assessment, SPPB,			higher prevalence of
	identify older CKD-			mini mental, clock			malnutrition (58 vs 29%, p =
	patients that may benefit			drawing			0.0005), physical impairment
	the most from a CGA.						(100% vs 78%; p < 0.0001),
							cognitive dysfunction (83% vs
							37%; p < 0.0001) and
							depression (50% vs 21%; p <
							0.001) compared to robust
							ones (NF-CKD). Moreover, F-
							CKD patients had higher
							probability to have > 2
							impaired domains (83%
							sensitivity and 76% specificity)
							respect to NF-CKD individuals.
							respect to MI -CND illumiduals.

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Vezza et al 2019	To retrospectively	Retrospective	Pre dialysis	Frailty index (FI)	N	Mortality and	66% of patients were frail
Italy [78]	calculate FI in a sample of					hospitalisation	(FI>0.25). Mean FI was 0.29
	patients with advanced						(standard deviation 0.10, range
	CKD. Its capacity to predict						0.08-0.66). The FI was
	hospitalizations, initiation						significantly correlated with
	of renal replacement						age and eGFR. The FI was
	therapy (RRT), and death.						predictive of hospitalization
							and mortality, even after
							adjustment for age and sex
Villareal et al 2014	To determine predictive	Cohort	Pre dialysis	Fried phenotype	Υ	Choice of	Of the 20 patients scheduled
Spain [79]	factors for the decision					conservative care	for CC, 18 (90%) were prefrail
	between HD or						versus 10 (28%) in the HD
	conservative care in						group (p = 0.000). Non-frail
	over75's						patients included 2 (10%) in
							the CC group versus 26 (72%)
							in the ITD group (p = 0.000). In
							multivariate analysis age and
							pre-frailty remained as
							independent predictors for the
							election of CC. During follow-
							up an increase of frail patients
							in both groups was observed.

Voorend et al	The primary aim was to	Qualitative	ESKD	> 65 years with a	Υ	N/A	Patients and professionals
Voorend et al 2021 Netherlands [80]	The primary aim was to elicit perspectives and experiences of patients and professionals with geriatric assessment (GA) in the care for older (≥65 years) patients approaching ESKD (estimated glomerular filtration rate < 20 ml/min/1.73m2), and to identify benefits, facilitators and barriers for implementation into routine nephrological care	Qualitative	ESKD	> 65 years with a geriatric impairment	Υ	N/A	Patients and professionals recognized increased vulnerability and (cognitive) comorbidity often unrelated to calendar age. Both believed patients need additional support in various geriatric domains. Patients regarded content/time spent on the GA predominantly positive. Professionals emphasised assessment increased awareness among the whole team for cognitive and social problems Outcomes of GA were enhanced dialogue on suitability of treatment options,(re)adjust treatment and provide/seek additional (social) support.
	routine nephrological care						were enhanced dialogue on suitability of treatment options,(re)adjust treatment and provide/seek additional

Voorend et al	The current study aimed	Pragmatic	Physical functioning is	Υ	Consensus of	The NGA contains instruments
2021 Netherlands	to reach agreement on a	consensus	assessed by Katz-ADL-		assessments and	in functional, cognitive,
[81]	nephrology tailored	based study	6, Lawton iADL,		measures to be	psychological, somatic, patient
	geriatric assessment		handgrip strength,		used in practice	preferences, nutritional status,
	(NGA) suitable to routinely		and falls		for NGA	and social domains. Selection
	identify major geriatric		questionnaire, 6CIT,			of instruments resulted from
	impairments in the target		GDS-15, LOT-R,			focus group meetings with
	population, which was		CFS,SF-12, Care giver			patients and professionals,
	defined as older patients		burden,			literature evidence, inventory
	(≥65 years of age) with		polypharmacy, SGA			of current geriatric screening
	stage G4–G5 CKD					practices, consensus between
						clinicians from nephrology and
						geriatrics, and pilot testing
Walker et al 2013	Cross sectional and	Systematic		N		The review identified 7 studies
Canada [82]	prospective studies in the	Review				associating frailty or physical
	general population and in					function to CKD. Of those, only
	the CKD population					two studies related frailty in
	examining the association					patients with CKD to a clinical
	between frailty and CKD					outcome. CKD was consistently
	and those relating frailty					associated with increasing
	in patients with CKD to					frailty or reduced physical
	clinical outcomes					function [odds ratios (OR) 1.30
						to 3.12]. In patients with CKD,
						frailty was associated with a
						greater than two-fold higher
						risk of dialysis and/or death
						[OR from 2.0 to 5.88

Walker et al 2015 Canada [83]	To determine the clinical history, prevalence of perceived and measured frailty and its association with dialysis treatment	Protocol CanFIT longitudinal observational study	Dialysis	Fried SPPB MoCA	N	Multiple Frailty Definitions: Short Physical Performance Battery (SPPB),	N/A
	choices and adverse outcomes in patients with advanced CKD					Fried Frailty Criteria, Frailty Index. Dialysis start: In-Centre Haemodialysis, Home Haemodialysis or Peritoneal Dialysis Outcomes: Death, Opt-out or Lost to	
						follow up	
Wu et al 2021 UK [84]	Our study aims to evaluate risk factors for mortality and the relative prognostic accuracy of various clinical assessment tools following hip fracture for patients living with CKD.	Secondary analysis of large prospective cohort study	CKD 3b-5	CFS CKD-F1, CI,KPS, DASI, Pre op risk	N	Mortality	The CFS had the best overall prognostic performance amongst the tools assessed. The CKD FI-LAB, CCI and KPS demonstrated good predictive accuracy for 30 day and 1 year mortality. AUC values for the pre-operative assessment scores, ASA index and NHFS were statistically signif cant for 30 day, but not for 1 year mortality
Wystma-Fisher et al 2021 [85]	The proposed Move More study will assess the feasibility of a physical activity intervention offered to the kidney failure inpatients	Protocol- intervention	CKD 5 and dialysis	Fried, GLTECQ,KDQOL, grip strength, sit to stand walking 15	N	The primary outcome of the study is the feasibility of administering an individualized early exercise/	N/A

						mobility intervention	
Yoneki et al 2019 Japan [86]	To assess the association between frailty and bone loss in patients undergoing haemodialysis	Cross sectional study	HD	Fried	N	Fried frailty measure, bone mass/Quantitative ultrasound calcaneal measurements	According to CHS criteria, 19 (21.1%) subjects were robust, 41 (45.6%) were pre-frail, and 30 (33.3%) were frail. ANOVA and chi-squared tests revealed that age, corrected Ca, and frailty components (with the exception of weight loss) significantly differed by frailty status. In this cross-sectional study of patients undergoing haemodialysis, frailty (according to CHS criteria) was significantly associated with calcaneal QUS parameters, including low SOS, BUA, and stiffness index for both sexes, even after adjusting for clinical characteristics. Interestingly, all QUS parameters declined significantly with increasing levels of frailty for both sexes.
Yoshida et al 2020 [87]	The aim of this study examined the correlation among CFS and other indices representing comorbidities, nutritional disorders, and geriatric syndrome, considering the prognosis.	Prospective cohort study	HD	CFS, SPICES score, CCI nutrition	N	Prognosis and morbidity	This study showed the relationship between CFS and CONUT score, CCI, and SPICES score in consideration of prognosis. As far as we know, there are no reports evaluating relationships of these indices and prognosis in the same patients with CKD. The CFS allows frailty to be defined and graded using simple clinical

Voung et al 2020	To evalure the persentions	Qualitative	HD	CFS	Yes	Outcomes	descriptors available from routine clinical assessment. The prognosis after initiation of dialysis is poor if the patient is frail during the preservation period. Important outcome identified
Young et al 2020 UK [88]	To explore the perceptions of frail HD patients in relation to participating in clinical research, IDC and a tailored exercise intervention.	component of main study	но		Yes	important to patients	form the qualitative data included animating mobility, maintaining activities of daily living and social support. Reporting of falls was not deemed important
Zanotto et al 2021 USA [89]	The objective of this study was to explore the diagnostic accuracy of several frailty screening methods, using the Fried phenotype as reference standard, in people receiving HD.	Cross sectional study	HD	Fried SF 36 TUG, STS-5, 15M Walk, IPAQ-SF	N	Accuracy of frailty assessments and Falls risk	The examined methods significantly discriminated frail from non-frail individuals, gait speed and TUG exhibiting the highest AUC values and elevated PPV/NPV. Gait speed had the highest specificity (93%) and PPV (0.86), TUG had the highest NPV (0.93). Accuracy of the same methods for fall-risk screening the Tinetti FES revealed the highest AUC value. While gait speed had an excellent diagnostic performance in people under 65 years of age, the TUG may be a more appropriate screening method for patients ≥ 65 years old). Importantly, the Tinetti FES was the only measure showing good diagnostic accuracy for both frailty and fall-risk screening.

Zhao et al 2020	Studies were considered	Systematic	HD	N	The prevalence of frailty range
China [90]	potentially eligible for	Review and			from 6.0% [5] to 82.0% [23]
	inclusion if they met the	meta-analysis			among studies and the pooled
	following criteria: (1) study				prevalence of frailty in patients
	designs observational or				on dialysis was 34.3% (95% Cl
	interventional providing				24.5–44.1%; z = 6.87; Age
	cross-sectional data on the				showed related to prevalence
	prevalence of frailty in				of frailty (β =0.54±0.19, t = 2.74,
	patients on haemodialysis				p = 0.018) p=0.00)
	based on any specific form				
	of frailty assessment, (2)				
	sample or subsample				
	consisted of patients on				
	haemodialysis. Studies				
	were excluded if they				
	were review, editorial,				
	comment, or conference				
	abstract				

Abbreviations used in the table (Full descriptor of assessments can be seen in appendix 2 of main paper)

ESRD/ESKD-End stage kidney disease	CFS -Clinical Frailty score	HGS-Hand grip strength
CKD -Chronic kidney disease	ADL-Activities of daily living	STS-Sit to stand
ACKD-Advanced chronic kidney disease	TUG-Timed up and go	DMMS - Dialysis Morbidity Mortality Study definition
HD-Haemodialysis	SPPB-Short physical performance battery	
PD - Peritoneal dialysis a= assistance	MoCA-Montreal cognitive assessment	
APD-Automated peritoneal dialysis a=assistance	IPAQ- International physical activity short form	
GA - Geriatric assessment	ISAR-H- Identification of seniors at risk of hospitalisation	
HHD- Home haemodialysis	F1-L-Frailty index lab	

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