

Adverse Health Effects of Frailty in CKD Patients

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Cause

	Effect (descriptions)	Risk Difference	CKD severity	Frailty Assessment	Sample Size	Reference
Lifestyle	Smoking	RR 1.18 (1.04-1.34)	CKD stage 5D (hemodialysis)	Fried Phenotypes	205	¹
Ethnicity	Hispanic	Frailty scores +0.6 points per year	CKD stage 5D (hemodialysis)	Fried Phenotypes	762	²
Biological						
Cardiovascular	Hypertension	RR 1.6 (1.26-2.04)	CKD stage 5D (hemodialysis)	Fried Phenotypes	205	¹
	Peripheral vascular disease	RR 1.58 (1.34-1.8)	CKD stage 5D (hemodialysis)	Fried Phenotypes	205	¹
	Left ventricular dysfunction	RR 1.18 (1.03-1.36)	CKD stage 5D (hemodialysis)	Fried Phenotypes	205	¹
	Endothelial dysfunction	$r = -0.367$ ($p = 0.004$) OR 3.86 (1.00-14.88)	CKD stages 3-5	Fried Phenotypes	61	³
Vascular access	Use of a permanent vascular access (fistula or graft)	HR 0.71 (0.51-0.98)	CKD stage 5D (hemodialysis)	Fried Phenotypes	2275	⁴
Cerebrovascular	Cerebrovascular Accident	RR 1.34 (1.19-1.5)	CKD stage 5D (hemodialysis)	Fried Phenotypes	205	¹
Pulmonary	COPD	OR 1.68 (1.16-2.45)	CKD stages 1-5	Fried Phenotypes	10256	⁵
Inflammatory	IL-6	Worse frailty	CKD stage 5D	Fried	762	²

已註解 [W使1]: 要寫一下是相對於什麼(ex. fistula/graft vs. catheter)

			(hemodialysis)	Phenotypes		
	CRP	After adjustment, OR	Chronic kidney	Fried	5888	⁶
	Fibrinogen	1.76 (1.28-2.41) to 1.50 (1.07-2.09)	insufficiency, serum creatinine \geq 1.3 mg/dL	Phenotypes		
Endocrinologic/ Metabolic	Diabetes	Frailty scores +0.7 points per year	CKD stage 5D (hemodialysis)	Fried	762	²
		OR 1.68 (1.16-2.45)	CKD stages 1-5	Phenotypes	10256	⁵
	Obesity (IMC \geq 30 kg/m ²)	OR 6.63 (1.16-36.77)	CKD stages 3-5	Fried	61	³
				Phenotypes		
	Higher parathyroid hormones (PTH)	r= 0.30 (p= 0.01)	CKD stages 3-5	Fried	61	³
				Phenotypes		
Body Composition	Higher fat mass	r= 0.25 (p= 0.04)	CKD stages 3-5	Fried	61	³
				Phenotypes		
Cancer	Cancer	OR 1.89 (1.19-2.99)	CKD stages 1-5	Fried	10256	⁵
				Phenotypes		
Arthritis	Arthritis	OR 3.34 (2.08-5.38)	CKD stages 1-5	Fried	10256	⁵
				Phenotypes		
Laboratory Data	eGFR (mL/min/1.72m ²) reference: > 60					
	eGFRcys < 30	Frailty prevalence 2.8	CKD stages 1-4	Fried	336	⁷
				Phenotypes		
	eGFRcys 30-44	Frailty prevalence 2.1				

Serum Albumin Concentrations (g/dL)	Frailty scores -1.1 points per g/dL	CKD stage 5D (hemodialysis)	Fried Phenotypes	762	²
Serum Creatinine <4 mg/dL*	RR 1.46 (1.22-1.71)	CKD stage 5D (hemodialysis)	Fried Phenotypes	205	¹
Testosterone, every 50% lower free form		CKD stage 5D (hemodialysis), men	Fried Phenotypes	440	⁸
being frail	OR 1.40 (1.05-1.53)				
becoming frail over 12 months	OR 1.40 (1.07-1.73)				
Hemoglobin	Adjusted, OR 1.76 (1.28-2.41) to 1.50	Chronic kidney insufficiency, serum creatinine ≥1.3mg/dL	Fried Phenotypes	5888	⁶
LDL, HDL	(1.07-2.09)				

Non-adjusted differences in comorbidity and system-based influences

	Effect (descriptions)	Prevalence	CKD Severity	Frailty Assessment	Sample Size	Reference
Biological						
Cardiovascular	Heart Failure (%)	30% vs 12%	CKD stages 1-4	Fried	336	7
	Angina (%)	34% vs. 22%	CKD stages 1-4	Fried	336	7
Cerebrovascular	Cerebrovascular Disease (%)	26.4 vs. 12.0	CKD stage 5D (hemodialysis)	Fried	324	9
Vascular Access	Catheter vs. Native/Prosthetic FAVI (%)	61.6% vs. 27.78/11.11%	CKD stage 5D (hemodialysis)	Fried	320	10
Endocrinologic/ Metabolic	Diabetes (%)	F vs. NF				
		64% vs. 49%	CKD stages 1-4	Fried	336	7
	Obesity (%)	F vs. NF				
		64% vs. 50%	CKD stages 1-4	Fried	336	7
		51.8% vs. 23.9%	CKD stage 5D (hemodialysis)	Fried	324	9
	BMI based on dry weight	31.5 vs. 27.6	CKD stage 5D (hemodialysis)	Fried	324	9
	BMI	22.53 vs. 26.16	CKD stage 5D	Fried	320	10

已註解 [W使2]: prevalent diseases

已註解 [W使3]: prevalent disease

已註解 [W使4]: characteristics of hemodialysis initiates
!! 當在沒有 cerebrovascular disease 的患者當中, frailty 跟 cognitive function 沒有什麼關係。

已註解 [W使5]: Characteristics
!! 好像比較少人提到

已註解 [W使6]: prevalent diseases

已註解 [W使7]: prevalent diseases

已註解 [W使8]: characteristics of dialysis initiates

已註解 [W使9]: characteristics of dialysis initiates

已註解 [W使10]: characteristics

			(hemodialysis)	Phenotypes		
Cognitive	Executive Function	F vs. NF				
	Trail Making Tests A (TMTA) scores	+12.08	CKD stage 5D (hemodialysis)	Fried Phenotypes	324	9
	Trail Making Tests B (TMTB) scores	+33.15	CKD stage 5D (hemodialysis)	Fried Phenotypes	324	9
Immunological	Inflammatory					
	CRP (ln CRP) (mg/dL)	1.12 vs 0.28	CKD stage 5D	Clinical Frailty Scale (CFS)	119	11
	IL6 (ln IL6) (mg/dL)	2.45 vs. 1.58	(peritoneal dialysis)			
	Mycophenolate mofetil (MMF) dose reduction (MDR)	F vs. NF	CKD stage 5T	Fried Phenotypes	525	12
	1 year since KT (%)	44 vs 40				
	2 years since KT (%)	54 vs. 45				
	3 years since KT (%)	67 vs. 51				
	Viral infection	F vs. NF				
Body Composition	Lower lean mass					
	Lean body mass (in frail vs. nonfrail) (%)	57.1% vs .14.7%	CKD stage 1-5	Edmonton Frail Scale (EFS)	41	13
Laboratory Data	Prealbumin (PRAB) (mg/dL)	28.9 vs. 38.3	CKD stage 1-5	Edmonton Frail Scale	41	11

已註解 [W使11]: at hemodialysis initiation

已註解 [W使12]: 跟 laboratory data 中的 PRAB (prealbumin), albumin 一樣都是以 frailty status 看不同

已註解 [W使13]: 在 complications 有 adjusted 的 data 了

已註解 [W使14]: 是 univariate 的，而且是分析 risk factor for frailty，我覺得要刪掉

已註解 [W使15]: 根據 frailty status 分的不同

(EFS)					
Serum albumin (mg/L)	Pearson correlation coefficient $r = -0.263$ ($p = 0.025$)	CKD stage 5D (hemodialysis)	Fried Phenotypes	74	¹⁵
	38 vs. 41	CKD stage 1-5	Edmonton Frail Scale (EFS)	41	¹³
	29.2 vs. 34.8	CKD stage 5D (peritoneal dialysis)	Clinical Frailty Scale (CFS)	119	¹¹
	36.1 vs. 38.5	CKD stage 5D (hemodialysis)	Fried Phenotypes	320	¹⁰
Frail with depression vs. Frail without depression vs. Nonfrail	32.9 vs. 34.9 vs. 35.8 ($p=0.025$)	CKD stage 5D (peritoneal dialysis)	In-house Chinese questionnaire	178	¹⁶
Calcium (mmol/L)	2.24 vs. 2.36	CKD stage 1-5	Edmonton Frail Scale (EFS)	41	¹³
Hemoglobin (g/dL)	Pearson correlation coefficient $r = -$	CKD stage 5D (hemodialysis)	Fried Phenotypes	74	¹⁵

已註解 [W使16]: 單位不同，請統整。兩位數的話應是 mg/L

已註解 [W使17]: 根據 frailty status 的不同

已註解 [W使18]: Creatinine (umol/L) , 299 vs. 115

		0.336 (p = 0.004)				
		6.49 vs. 8.07 (p = 0.037)				
		10.35 vs. 10.97	CKD stage 5D (hemodialysis)	Fried Phenotypes	320	10
	25-hydroxy vitamin D (ng/mL)	Pearson correlation coefficient r = -0.363 (p = 0.002)	CKD stage 5D (hemodialysis)	Fried Phenotypes	74	15
		11.58 vs. 17.09 (p = 0.005)				
Miscellaneous	Dialysis clearance rate (Kt/V and urea reduction ratio)	↑	CKD stage 5D (chronic dialysis)	FRAIL scale	46	17
Composite	Number of complications (complications identified at data collection: High Pressure Cramping, Anemia, Weight loss Pain, Weakness, Weight gain Constipation, Heart Arrhythmia, Headache, Itch, Recurrent infections, Arterial hypertension)	Spearman's correlation 0.666 (p = 0.000 in table) (p < 0.05 in text)	Elderly (≥ 60 yo), with diagnosis of CKD	Edmonton Frail Scale (EFS)	35	18
	Higher number of comorbid conditions	6 vs. 4 (p = 0.03)	CKD stage 1-5	Edmonton Frail Scale	41	13

已註解 [W使19]: Clearance rate 哪一種？

				(EFS)		
	Charlson's comorbidity score	Spearman's rank correlation coefficient $r = 0.40$ ($p < 0.0001$)	CKD stage 5D (peritoneal dialysis)	Chinese questionnaire	193	¹⁹
Functional Status	Disability	F vs. NF				
	≥1 disability in activities of daily Living (ADLs)	15% vs. 5%	CKD stages 1-4	Fried Phenotypes	336	⁷
	≥1 disability in instrumental activities of daily living (IADLs)	60% vs. 28%				
	≥1 disability in mobility tasks	40% vs. 18%				
	Less ADL-independent	55% vs. 91%	≥ 65 yo, predialysis, eGFR < 20 mL/min	Groningen frailty indicator (GFI)	65	²⁰
	Ability to perform basic activities of daily living	33.33% vs 76.4%	CKD stage 5D (hemodialysis)	Fried Phenotypes	320	¹⁰
	Ability to perform transfers	38.8% vs. 84.7%				
Microbiota						

已註解 [W使20]:

已註解 [W使21]: Mentioned in other articles as complications, can omit

Gut microbiota	Malnutrition-Inflammation-Score (MIS)	7.6 vs. 3.9	Stage 3b-4, eGFR 15-45ml/min	Fried Phenotype score	64 (and 15 control subjects)	21
	Abundance of unclassified Mogibacteriaceae and Oscillospira	Directly proportional to MIS				
	Abundance of Akkermansia, Ruminococcus, and Eubacterium	Inversely proportional to MIS				
	Bacterial Abundance of some genera (Mogibacteriaceae, Coriobacteriaceae, Eggerthella, Erwinia, Coprobacillus, Anaerotruncus, etc)	↑				
Neurological						
EEG	Brain Wave	F vs. NF	ESRD, under chronic dialysis	FRAIL scale	46	17
	Global DAR	283 ± 679 vs. 2971 ± 4859				
	DARs (left frontal)	135 ± 250 vs. 3073 ± 4702				
	DAR (left TO)	197 ± 318 vs. 3708 ± 6398				
	DAR (central)	55 ± 96 vs. 1773 ± 3262				
	DAR (right TO)	187 ± 261 vs.				

		4400 ± 7763				
Global DTABR		191 ± 469 vs. 1781 ± 2793				
DTABR (left frontal)		86 ± 158 vs. 1680 ± 2388				
DTABR (left TO)		130 ± 210 vs. 1884 ± 2828				
DTABR (central)		39 ± 65 vs. 1132 ± 1957				
DTABR (right TO)		126 ± 178 vs. 2960 ± 5271				
Psychological						
Mood	Mood Change	Negative change	CKD stage 5D (hemodialysis)	Edmonton Frail Scale (EFS)	N/A	22
Mental Health						
Anxiety	Hospital Anxiety and Depression Scale (HADS)	<u>Women</u> : ↑ in global, psychological, social frailty <u>Men</u> : ↑ in Physical frailty	ESRD, under online-haemodiafiltration (OL-HDF)	N/A	97	23
Depression	Hospital Anxiety and Depression Scale (HADS)	<u>Men</u> ↑ in global,	ESRD, under online-	N/A	97	23

		psychological, physical frailty	haemodiafiltration (OL-HDF)			
	Incidence (%) (Self-reported Major Depression Inventory)	83 vs. 6	CKD stage 1-5	Edmonton Frail Scale (EFS)	41	¹³
Mental Function	Post-KT delirium	9.0% vs. 3.9%	CKD stage 5T	Fried Phenotypes	893	²⁴
Sociological						
Interaction	Interaction with family	Good				²⁵
Physical activity	Minnesota Leisure Time Activity (LTA)	95 vs. 735 (p<0.001)	CKD stage 5D (hemodialysis)	Fried Phenotypes	68	²⁶
	Low Physical Activity Questionnaire (LoPAQ)	280 vs. 798 (p=0.003)				
	Sitting (hours/day)	6.5 vs. 5 (p=0.04)				
Nutritional Status	Overall subjective global assessment (SGA) (weight loss, anorexia, subcutaneous fat, muscle mass) (Frail with depression vs. Frail without depression vs. Nonfrail)	5.04 vs. 5.41 vs. 5.75 (p < 0.0001)	CKD stage 5D (peritoneal dialysis)	In-house Chinese questionnaire	178	¹⁶
		Spearman's rank correlation coefficient r = - 0.44, p < 0.0001	CKD stage 5D (peritoneal dialysis)	Chinese questionnaire	193	¹⁹

	Malnutrition inflammation score (MIS) (frail with depression vs. frail without depression vs. nonfrail)	9.48 vs. 7.13 vs. 5.12 (p < 0.0001)	CKD stage 5D (peritoneal dialysis)	In-house Chinese questionnaire	178	¹⁶
		Spearman's rank correlation coefficient r = 0.40, p < 0.0001	CKD stage 5D (peritoneal dialysis)	Chinese questionnaire	193	¹⁹
Quality of Life	HRQoL					
	SF-36					
	Scores in physical functioning, blood pressure, role physical, and physical component summary domains	↓	CKD stage 1-5	Edmonton Frail Scale (EFS)	41	¹³
	Kidney Disease Quality of Life (KDQoL)					
	Physical health	33.7 vs. 40.7	ESRD CKD stage 5D (conventional hemodialysis)	Fried Phenotypes	151	²⁷
	Kidney disease effects	51.6 vs. 66.8				
Independence	Functional Independence Measure (FIM)		Elderly, ≥65y/o	Edmonton	137	²⁸
	Spearman's correlation	Frailty diagnosis with global FIM -0.703 (p<0.001)		Frail Scale (EFS)		

	coefficient	Frailty diagnosis with motor FIM	-0.714 (p<0.001)			
		Frailty diagnosis with cognitive FIM	-0.575 (p<0.001)			
		EFS scores with gross FIM	-0.53 (p<0.01)			
Health-care utilization	Hospitalization		90% vs. 53% (p = 0.04)	≥ 65 yo, predialysis, eGFR < 20 mL/min	Groningen frailty indicator (GFI)	65 ²⁰
		Cumulative number of inpatient health-care visits	↑	CKD stage 1-5	Edmonton Frail Scale (EFS)	41 ¹³
		Cumulative number of emergency health-care visits				
		Cumulative number of total health-care visits				
		Admissions/year	0.77727 vs. 0.2838	CKD stage 5D (hemodialysis)	Fried Phenotypes	320 ¹⁰

Complications

	Effect (descriptions)	Risk Difference	CKD Severity	Frailty Assessment	Sample Size	Reference
Biological						
Cardiovascular	Cerebrovascular Accident	OR 1.55 (1.05-2.99)	CKD stage 5D (hemodialysis)	Fried Phenotypes	2275	⁴
	QRS duration	β coefficient = -0.29, t = -2.03 (p = 0.048)	CKD stage 5D (chronic hemodialysis)	Edmonton frailty scale (EFS)	41	²⁹
		β coefficient = -0.27, t = -1.84 (p = 0.05)		FRAIL scale		
	Vascular Access failure	HR 2.63 (1.03-6.71)	ESRD (CKD stage 5D)	Self-reported FRAIL scale	51	³⁰
Renal Function Decline	Risk for death or dialysis therapy	2.5 (1.4-4.4)-fold greater	CKD stages 1-4	Fried Phenotypes	336	⁷
Immunological	Mycophenolate mofetil (MMF) dose reduction (MDR)	HR 1.29 (1.01-1.66)	CKD stage 5T	Fried Phenotypes	525	¹²
Cognitive	Modified Mini-Mental State (3MS)	-2.37 to -2.80 (1 year) (p=0.03)	ESRD	Fried Phenotypes	324	⁹
	Declined, 1-4 years post-KT (points/week)	Slope = -0.04 vs. 0.005	CKD stage 5T	Fried physical frailty phenotypes (PFP)	665	³¹
	At 4 year post-KT (points)	-5.5 (87.4 vs.				

已註解 [W使22]: simple

已註解 [W使23]: simple

		92.9)				
Functional status	Disability	F vs. Prefrail vs. NF				
	Need for activities of daily livings (ADL) assistance	OR 11.32 (5.49-23.32) vs. 1.93 (1.01-3.68) vs. 1.00	CKD stage 5D (hemodialysis)	Fried Phenotypes	742	³²
Diabetes	Diabetes	OR 1.35 (1.10-1.65)	CKD stage 5D	Fried Phenotypes	2275	⁴
Body composition	Lower lean mass	F/PF vs. NF				
	Whole body (kg)	34.7 vs. 43.1	CKD stage 5D (hemodialysis)	Self- report instrument evaluating five dimensions of frailty (fatigue, resistance, ambulation, illnesses, and weight loss)	44	¹⁴
	Cephalic/trunk (g)	3059 vs. 3288 (cephalic); 17.4 vs. 22.1 (trunk)				
	Right/Left upper limb (g)	1831 vs. 2493 (right); 1869 vs. 2515 (left)				
	Right/Left lower limb (g)	4920 vs. 6114 (right); 4650 vs. 6349 (left)				
	Bones					

Bone Mineral Density (BMD)	One year follow-up, with frailty	ESRD	FRAIL Scale	43	33
L1	$\beta = -0.4$, t = -2.18, p=0.04	CKD stage 5D (chronic hemodialysis)			
L4	$\beta = -0.39$, t = -2.1, p=0.046				
Femur Neck (FN)	$\beta = -0.5$, t = -2.96, p<0.01				
Total	$\beta = -0.53$, t = -3.27, p < 0.01	ESRD	FRAIL Scale	43	33
		CKD stage 5D (chronic hemodialysis)			
Areas	One year follow-up, with frailty	ESRD	FRAIL Scale	43	33
Average L-spine areas	$\beta = -0.48$, t = -2.84, p < 0.01	CKD stage 5D (chronic hemodialysis)			
Changes of average L-spine areas	$\beta = -0.5$, t = -3.02, p<0.01				
Z-score	One year follow-up, with frailty				

Percentage change of L1 Z-score	$\beta = -0.45$, t = -2.11, p=0.049				
Vertebral Compression Fracture (VCF)	OR 1.8 (p = 0.01)	ESRD CKD stage 5D (chronic hemodialysis)	FRAIL Scale	43	34
Quantitative ultrasound (QUS) parameters of calcaneus					
Speed of sound (SOS)	Standardized β (range, p value)	CKD stage 5D (maintenance hemodialysis)	Fried Phenotypes	214	35
Female (Five frailty criteria)	Negative (-0.253 to -0.439, p \leq 0.034)				
Male (All criteria significant except weight loss)	Negative (-0.277 to -0.402, p \leq 0.003)				
Broadband ultrasound attenuation (BUA)	Standardized β (range, p value)				
Female (All criteria significant except weakness and weight loss)	Negative (-0.209 to -0.354, p \leq 0.045)				

Male (All criteria significant except weight loss)	Negative (-0.171 to -0.371, $p \leq$ 0.045)				
Stiffness index	Standardized β (range, p value)				
Female (All criteria significant except weight loss)	Negative (-0.271 to -0.461, $p \leq$ 0.018)				
Male (Five frailty criteria)	Negative (-0.183 to -0.461, $p \leq$ 0.048)				
Muscles					
Quadriceps muscle area (magnitude of association with PbF vs. 10 years of age)	Multivariable coefficient -30.3 cm^2 ($p = 0.02$) vs. -6.6 cm^2 ($p =$ 0.0001)	CKD stage 5D (hemodialysis)	Performance-based frailty (PbF)	80	36
Appendicular skeletal muscle mass index (ASMI)	Unadjusted 6.8 vs. 7.7; adjusted $p < 0.05$	CKD stage 1-5	Edmonton Frail Scale (EFS)	41	13
Fat					

	Appendicular fat percentage	FRAIL scale				
	Left/Right lower extremity	t = 2.32; p = 0.03 (left); t = 2.05; p = 0.048 (right)	CKD stage 5D (hemodialysis)	FRAIL scale	44	14
	Left/Right upper extremity	t = 2.66; p = 0.01 (left); t = 3.09; p = <0.01 (right)				
	Appendicular fat percentage	Frail/Prefrail vs. Nonfrail				
	Left/Right lower extremity	t = 2.31; p = 0.03 (left); t = 2.28; p = 0.03 (right)	CKD stage 5D (hemodialysis)	Self- report instrument evaluating five dimensions of frailty (fatigue, resistance, ambulation, illnesses, and weight loss)	44	14
	Right upper extremity	t = 2.35; p = 0.03				
	Sarcopenia	aOR 12.2 (2.27-65.5)	CKD stage 5D (peritoneal dialysis)	Clinical Frailty Scale (CFS)	11912	11
Laboratory data	Serum Albumin Concentrations (g/dL)					
	<3.2 vs. ≥3.9	OR 1.89 (1.30-2.59)	CKD stage 5D	Fried Phenotypes	2275	4
	Hypoalbuminemia	Negative association (p =	CKD stage 5D (maintenance	Frail Scale	46	37

已註解 [W使24]: 是 multivariate linear regression

已註解 [W使25]: adjusted for blah, blah, blah, risk factor for sarcopenia

		0.01)	hemodialysis) (ESRD)			
Psychological						
Delirium	Post-KT delirium	OR 2.05 (1.02-4.13)	CKD stage 5T	Fried Phenotypes	893	²⁴
Distress	Distress Thermometer	$\beta = 0.35$, $t = 3.0$ (95% CL = 0.12-0.58) ($p = 0.003$)	CKD stage 5D (hemodialysis)	Canadian frailty score	382	³⁸
Quality of Life						
	HRQoL					
	Fair/Poor HRQOL at follow-up (median 9.4 mo)	aOR 2.79 (1.32-5.90)	ESRD CKD stage 5T	Fried Phenotypes	233	³⁹
	Worsening HRQOL at follow-up (median 9.4 mo)	aRR 2.91 (1.08-7.80)				
	SF-36					
	Hierarchical regression R ² change (effects of frailty on HRQoL) in Physical Component Summary (PCS)	29% ($p < 0.001$)	CKD stage 2-4	Fried Phenotypes	168	⁴⁰
	Hierarchical regression R ² change (effects of frailty on HRQoL) in Mental Component Summary (MCS)	21.3% ($p < 0.001$)				

Physical components	Simple linear regression coefficient = -1.12 (-1.47 to -0.76) (p < 0.001)	CKD stages 3-5 (predialysis treatment)	Fried Phenotypes	61	41
Mental components	Simple linear regression coefficient = -0.75 (-1.40 to -.016)				
SF-12					
MCS	Effect estimate 0.94 (0.91-0.97) (p<0.01)	CKD stage 5D (peritoneal dialysis, n=129; hemodialysis, n=122)	The Canadian Study of Health and Aging Clinical Frailty Scale (CFS)	251	42
PCS	Effect estimate 0.88 (0.84-0.91) (p<0.01)				
KDQOL-SF scores within 3 months post-KT					
At KT	↓	CKD stage 5T	Fried Phenotypes	443	43
Physical HRQoL	-6.31 points (95% CI -8.16 to -4.46)				
Kidney disease-	-6.53 points				

specific HRQoL	(95% CI -9.17 to -3.89)
Post-KT	Greater improvement
Physical HRQoL	1.35 points/month (0.65 to 2.05) vs. 0.34 points/month (-0.17 to 0.85)
Kidney disease-specific HRQoL	3.75 points/month (2.89 to 4.60) vs. 2.41 points/month (1.78 to 3.04)
Constituent domains	Greater improvement
General health	4.93 points/month (3.51 to 6.35) vs. 2.87 points/month (1.82 to 3.92)

Effects of ESRD on daily living	7.10 points/month (5.68 to 8.51) vs. 4.01 points/month (2.99 to 5.03)				
Cognitive function	2.88 points/month (1.80 to 3.96) vs. 1.28 points/month (0.50 to 2.07)				
Social interaction	1.18 points/month (-0.06 to 2.43) vs. -0.57 points/month (-1.47 to 0.33)				
Illness Intrusiveness Rating Scale	Effect estimate 1.14 (1.09-1.20)	CKD stage 5D (peritoneal dialysis, n=129; hemodialysis, n=122)	The Canadian Study of Health and Aging Clinical Frailty Scale (CFS)	251	42
Barthel Index	Effect estimate 0.89 (0.86-.093)				
Symptom score	Effect estimate 1.23 (1.13-1.34)				

	Hospital Anxiety and Depression Scale	Effect estimate 1.21 (1.11-1.31)				
	Falls	HR 2.1 (1.21-3.92)	CKD stage 5D (hemodialysis)	Fried Phenotypes	205	¹
		OR 2.39 (1.22-4.71)	CKD stage 5D (maintenance hemodialysis)	Fried frailty index	762	⁴⁴
	Higher numbers of falls	HR 3.09 (1.38-6.90)	CKD stage 5D (hemodialysis)		95	⁴⁵
	Time to first fall or fracture requiring medical attention	HR 1.60 (1.16-2.20)	CKD stage 5D (maintenance hemodialysis)	Modified Fried Phenotypes by Bao Y ⁴⁶ .	1646	⁴⁷
Graft Loss	Death-censored graft loss					
	F vs. NF (in patients with depressive symptoms)	aHR 6.20 (1.67, 22.95) vs. 3.16 (0.90, 11.04)	CKD stage 5T	Fried Phenotypes	773	⁴⁸
Health-care utilization	Hospitalization/Death	HR 1.56 (1.36-1.79)	CKD stage 5D	Fried Phenotypes	2275	⁴
	Hospitalization	HR 2.06 (1.18-3.58)	CKD stage 5D (hemodialysis)	Fried Phenotypes	205	¹
		aHR 1.80 (1.4-2.3)	CKD stage 5D (maintenance hemodialysis &	Adopted	1658	⁴⁹

		peritoneal dialysis)			
	Relative risk = 1.43 (1.00-2.03)	CKD stage 5D (hemodialysis)	Fried Phenotypes	146	⁵⁰
Number of hospitalizations for all causes	beta = 0.29 (p < 0.0001)	CKD stage 5D (peritoneal dialysis)	Chinese questionnaire	193	¹⁹
Number of hospitalizations related to cardiovascular events	beta = 0.37 (p < 0.0001)				
Non-vascular access- related hospitalizations	aHR 1.98 (1.41- 1.87) (內文應該 寫錯，無勘誤)	CKD stage 5D	Fried Phenotypes	2275	⁴
Time to first hospitalization	HR 1.26 (1.09- 1.45)	CKD stage 5D (maintenance dialysis)	Earlier modification of Fried Phenotypes by Johansen et al ⁴ , but without weight loss.	1576	⁴⁶
Early Hospital Readmission (EHR)	aRR 1.61 (1.81- 2.19) (p=0.002)	CKD stage 5T	Fried Phenotypes	383	⁵¹
Duration of hospitalization					
Hospital stay (days per year of follow up) (frail with depression vs. frail without depression vs. nonfrail)	26.62 (IQR 10.65-61.18) vs. 14.05 (IQR 3.57- 37.27) vs. 8.04 (IQR 0.91-19.42) (p<0.0001)	CKD stage 5D (peritoneal dialysis)	In-house Chinese questionnaire	178	¹⁶

	Total length of hospital stay	beta = 0.34 (p < 0.0001)	CKD stage 5D (peritoneal dialysis)	Chinese questionnaire	193	¹⁹
	Longer Length of Stay (LOS)					
	with delayed graft function (DGF), LOS	Relative Risk 1.15 (1.03-1.29)	CKD stage 5T	Fried Phenotypes	589	⁵²
	With DGF, LOS ≥2 weeks	OR 1.57 (1.06-2.33)				
	≥2 weeks		CKD stage 5 to 5T	Fried Phenotypes	569	⁵³
	Change in 3 categories (more frail)	OR 2.02 (1.20-3.40)				
	Change in frailty scores (more frail)	OR 1.92 (1.13-3.25)				
	With depressive symptoms (aRR difference between F and NF)	aRR 1.88 (1.70-2.08) vs. 1.38 (1.27-1.52)	CKD stage 5T	Fried Phenotypes	773	⁴⁸
	CES-D score (10-point increase) (aRR increase between F and NF)	aRR 1.23 (1.16-1.31) vs. 1.17 (1.08-1.27)				
Mortality	Mortality	2.17 fold	CKD stage 5T	Fried Phenotypes	537	⁵⁴
		HR 1.57 (1.25-1.97)	CKD stage 5D (maintenance dialysis)	Earlier modification of Fried Phenotypes by Johansen et al ⁴ , but without weight loss.	1576	⁴⁶
		HR 2.24 (1.60-	CKD stage 5D	Fried Phenotypes	2275	⁴

	3.15)				
	HR 1.22 (1.04-1.43)	CKD stage 5D (incident chronic dialysis)	CFS	390	⁵⁵
	HR 4.28 (1.22-14.98)	Predialysis (eGFR \leq 25 mL)	PRISMA questionnaire & Timed up and Go test	104	⁵⁶
	aHR 9.83 (1.80-53.7)	CKD stage 5D (peritoneal dialysis)	CFS	119	¹¹
	HR 2.60 (1.04-6.49)	CKD stage 5D (hemodialysis)	Fried Phenotypes	146	⁵⁰
	HR 2.37 (1.11-5.02)	CKD stage 5D (maintenance hemodialysis & peritoneal dialysis)	Adopted	1658	⁴⁹
	20.45% vs. 12.36% (p<0.005)	CKD stage 5D (hemodialysis)	Fried Phenotypes	320	¹⁰
Performance-based frailty	HR 2.16 (1.41-3.29)	CKD stage 5D (hemodialysis)	Fried Phenotypes & a definition that substitutes self-reported measures available on the Medical Outcomes Study 36-Item Short Form (SF-36) for the physical performance and	771	⁵⁷
Self-reported function-based frailty	HR 1.93 (1.24-3.00)				
Patients who met both performance- and self-reported function-based	HR 2.46 (1.51-4.01)				

frailty	exhaustion criteria.				
F vs. NF (in patients with depressive symptoms)	aHR 2.62 (1.03, 6.70) vs. 1.92 (0.68, 5.38)	CKD stage 5T	Fried Phenotypes	773	48
At 24-month follow up, frail with depression vs. frail without depression vs. nonfrail	62.5% vs. 71.4% vs 86.6% (p=0.001)	CKD stage 5D (peritoneal dialysis)	In-house Chinese questionnaire	178	16
Prediction ability of comorbidities in F vs. NF	HR 0.75 (0.44-1.29) vs. 1.66 (1.17-2.35)	CKD stage 5T (KT candidates, on waitlist)	Fried Phenotypes	2086	58
Out of 10 deceased within 1 year of initiation (percentage of F vs. NF)	30% vs. 9%	≥ 65 yo, predialysis, eGFR < 20 mL/min	Groningen frailty indicator (GFI)	65	20
Risk for death or dialysis therapy	2.5 (1.4-4.4)-fold greater	CKD stages 1-4	Fried Phenotypes	336	7
All-cause mortality					
Adjusted	HR 1.66 (1.03-2.67)	CKD stage 5D (incident chronic dialysis)	Fried Phenotypes	370	59
Among BMI ≥30 kg/m ²	HR 3.77 (1.10-12.92)				
Above median Waist-Hip Ratio (WHR)	HR 2.38 (1.17-4.82)				
Anti-neutrophil cytoplasmic	HR 2.43 (1.48-	CKD stage 5D to 5T	Inability to walk without	425	60

	antibody (ANCA)-associated vasculitides (AAV) patients' mortality	3.99)	(RRT [hemodialysis, peritoneal dialysis, transplantation])	help		
	Mortality of patients with monoclonal gammopathy and ESRD caused by myeloma cast nephropathy (MCN), immunoglobulin light chain amyloidosis (ALA), or light-chain deposition disease (LCDD)	HR, 1.93 (1.58-2.36)	CKD stage 5D to 5T (RRT [hemodialysis, peritoneal dialysis, transplantation])	Inability to walk without help	1462	⁶¹
	Post-KT mortality					
	Change in 3 categories (more frail)	HR 2.27 (1.11-4.65)	CKD stage 5 to 5T	Fried Phenotypes	569	⁵³
	Change in frailty scores (more frail)	HR 2.36 (1.12-4.99)				
Composite	Composite outcomes of all-cause death or cardiovascular hospitalization	HR 23.58 (1.61-346.03)	CKD stage 5D (hemodialysis)	Multidimensional frailty score based on comprehensive geriatric assessment (CGA) protocol	46	⁶²
	30-day postoperative (KT) complications according to Comprehensive Complication	β =13.31, 95% CI 5.72-20.89 (p = 0.0007)	CKD stage 5T	Groningen Frailty Indicator	150	⁶³

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