

Management of early chronic kidney disease

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Introduction

- A growing public health problem in NZ and throughout the world.
- Unknown prevalence in NZ, but 7 – 10% based on overseas population-based studies.
- Maori and Pacific peoples have higher rates of diabetes, CKD and ESKD in NZ.
- 2674 dialysis patients in 2015 (12% increase in the last 5 years).



Introduction

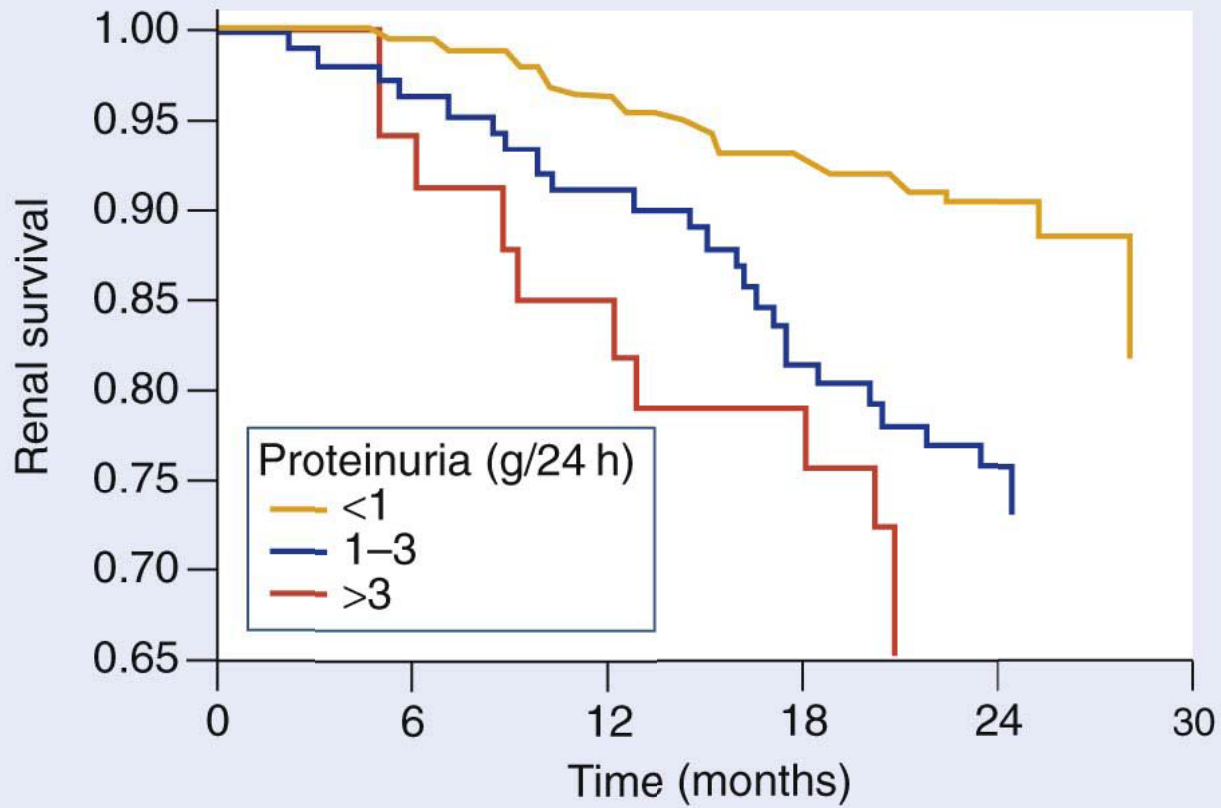
- A major risk factor for cardiovascular disease (CVD) and premature death.
- Timely detection and management of CKD reduce the risks of CVD and CKD progression by up to 50%.

The Classification of CKD and Prognostic Risk from the KDIGO CKD Consensus Consortium

Prognosis of CKD by GFR and Albuminuria Categories: KDIGO 2012 (units changed to SI)				Persistent albuminuria categories		
				Description and range		
				A1	A2	A3
				Normal to mildly increased < 3 mg/mmol	Moderately increased 3–30 mg/mmol	Severely increased > 30 mg/mmol
GFR categories (ml/min/1.73 m ²) Description and range	G1	Normal or high	≥ 90			
	G2	Mildly decreased	60–89			
	G3a	Mildly to moderately decreased	45–59			
	G3b	Moderately to severely decreased	30–44			
	G4	Severely decreased	15–29			
	G5	Kidney failure	< 15			

Green: low risk (if no other markers of kidney disease, no CKD); Yellow: moderately increased risk; Orange: high risk; Red: very high risk.

Renal Survival and Level of Proteinuria



Risk of CKD

All-cause mortality

	ACR < 1	ACR 1-3	ACR 3-30	ACR ≥30
eGFR > 105	1.1	1.5	2.2	5.0
eGFR 90-105	Ref	1.4	1.5	3.1
eGFR 75-90	1.0	1.3	1.7	2.3
eGFR 60-75	1.0	1.4	1.8	2.7
eGFR 45-60	1.3	1.7	2.2	3.6
eGFR 30-45	1.9	2.3	3.3	4.9
eGFR 15-30	5.3	3.6	4.7	6.6

Cardiovascular mortality

	ACR < 1	ACR 1-3	ACR 3-30	ACR ≥30
eGFR > 105	0.9	1.3	2.3	2.1
eGFR 90-105	Ref	1.5	1.7	3.7
eGFR 75-90	1.0	1.3	1.6	3.7
eGFR 60-75	1.1	1.4	2.0	4.1
eGFR 45-60	1.5	2.2	2.8	4.3
eGFR 30-45	2.2	2.7	3.4	5.2
eGFR 15-30	14	7.9	4.8	8.1

Screening for early CKD in primary care

- Population screening in isolation is not recommended.
- Major risk factors for CKD in NZ:
 - Hypertension
 - Diabetes
 - Age over 60 years
 - BMI > 35
 - Family history of CKD
 - Maori and Pacific ethnicity
 - CVD resulting in reduced renal perfusion and endothelial dysfunction
 - Prostatic syndrome / urologic disease which has the potential to cause obstructive uropathy

Screening for early CKD in primary care

- Other high-risk groups may include:
 - Previous acute kidney injury
 - Cigarette smoking
 - Nephrotoxic drugs
 - Systemic autoimmune conditions
 - Renal stones, recurrent urinary tract infections
- Frequency of testing – every 12 months
 - Urine ACR
 - Serum creatinine and eGFR

Firstly, repeat 1 – 2x
over 3 months, then

Urine PCR
MSU (haematuria)

Inaccurate in some patients groups:

- Children
- Abnormally low or high muscle mass
- Cirrhosis

Primary prevention of CKD

- Avoid development of CKD → a preferable strategy
- Limited evidence in general, but probably good evidence on the following:
 - Achieve blood pressure target of $< 140 / 90$ mmHg***
 - Educational programs
 - Avoidance or cessation of cigarette smoking
 - An individualized care plan with appropriate prescription of medications and interventions targeting CVD and renal risk modifications
- Importance of addressing and modifying risk factors irrespective of evidence

Managing patients with CKD in Primary Care

- Stable CKD stage 3 or those aged > 75 years with early and stable CKD stage 4 can be managed in primary care.
- The most important aspects of CKD management are:
 - Controlling blood pressure
 - Controlling blood glucose (if the patient has diabetes)
- Appropriate cardiovascular disease management.
- Complementary community-based care strategies involving nurse-led teams (the DEFEND trial).

Managing patients with CKD in Primary Care

Lifestyle management	Systolic blood pressure reduction
Reduce BMI to < 30, an ideal target of ≤ 25 Waist circumference < 102 cm - male Waist circumference < 88 cm - female	5 – 20 mmHg
Moderate intensity physical activity > 30min/day	4 – 9 mmHg
Low salt diet (< 6 g / day), avoid processed food and takeaway	2 – 8 mmHg
Reduce alcohol consumption ≤ 2 standard drinks / day – female ≤ 3 standard drinks / day – male	2 – 4 mmHg
Smoking cessation	-
Low sugar and cholesterol diet	-
Normal daily protein intake 0.75 – 1 g/kg/day	-
Adequate fluid intake 30ml/kg/day ***	-

Managing patients with CKD in Primary Care

- **Blood pressure management**

- Target blood pressure $\leq 130 / 80$ mmHg ***
- ACEIs or ARBs are the first line treatment
- Avoid the combination of ACEI and ARB
- Calcium channel blockers, thiazide and B-blockers are second line therapy

- **Glycaemic control**

- Target HbA1C < 53 mmol/mol ***
- Metformin can be used in stage 4 CKD, but maximal dose 500mg – 1g /day ***

Managing patients with CKD in Primary Care

- **Treating hyperlipidaemia according to cardiovascular risk**
 - Statin treatment, less effective in advanced CKD
 - Avoid Fibrates
- **Hyperuricemic and gout control**
 - Target uric acid < 0.36 mmol/L
 - Allopurinol is the first line therapy
 - Dosage up to 300mg/day in advanced CKD
 - Slow titration
 - Prednisone for acute gout attacks; Colchicine use with caution
 - Avoid NSAIDs

Managing patients with CKD in Primary Care

- **Avoid nephrotoxic agents**

- NSAIDs
- Proton pump inhibitors
- Lithium
- Antivirals (Acyclovir, Cidofovir, Foscarnet, Indinavir)
- Amphotericin
- Aminoglycosides (Amikacin, gentamicin, tobramycin)
- Calcineurin inhibitors (Ciclosporin/Tacrolimus)
- Chemotherapeutics (Cisplatin, Ifosfamide)
- Sulphonamides
- Fibrates
- Radiocontrast media

Managing patients with CKD in Primary Care

- **Avoid nephrotoxic agents**
 - Unregulated traditional/Herbal medicine
 - Heavy metals contamination
 - Aristolochic acid
- **Preventing acute kidney injury**
 - Avoid some or all of antihypertensive medications, diuretics, NSAID, metformin during an acute illness
 - Pre-hydration when undergoing procedures requiring radiocontrast media.

Monitoring patients with established CKD

CKD staging	Frequency of review	Investigations requested
Stage 1 – 2	6 – 12 months; less frequently if the patient's eGFR is stable and risk factors controlled	Serum creatinine, ACR (or PCR), serum electrolytes, serum urate, HbA _{1c} and lipids
Stage 3	Three to six-monthly	In addition to the above: FBC, serum ferritin, calcium, phosphate and parathyroid hormone
Stage 4	Three-monthly	In addition to the above: plasma bicarbonate
Stage 5	Monthly	Investigations usually determined in conjunction with a nephrologist

When to refer for specialist renal care

- Stage 4 and 5 CKD of any cause (eGFR < 30 ml/min/1.73m²)
- A progressive decline in eGFR from a baseline of < 60ml/min/1.73m²
 - > 5ml/min over a 6-month period, confirmed on >= 3 separate readings
- Evidence of intrinsic renal disease (GN, AIN, polycystic kidney disease)
- **Persistent** significant albuminuria (uACR > 30mg/mmol) or proteinuria (uPCR > 50mg/mmol) or urinary protein excretion >= 500mg/24h) and/or glomerular haematuria
- CKD and uncontrolled hypertension despite >= 3 BP medications

Take home message

- CKD is a major health burden in NZ.
- Early detection of CKD improves CVD outcome and slows CKD progression.
- The most important aspects of CKD management are:
 - Controlling blood pressure
 - Controlling blood glucose (if the patient has diabetes)
- Lifestyle modifications can reduce rate of renal function decline.
- Primary care physicians play a significant role in the management of early CKD

Reference

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