# Overview of CKD & Proteinuria Management in Primary Care

Dr Jay Hingwala December 11, 2020

# Faculty/Presenter Disclosure

- Faculty: Dr. Jay Hingwala
- Relationships with commercial interests:
  - Advisory Board: Otsuka and Janssen

# Mitigating Potential Bias

- All information presented will be supported by national and provincial guidelines
- No specific medications or trade names will be used

# Objectives

At the conclusion of this educational activity, participants should be able to:

- Appreciate the growing rate of CKD and ESRD in Manitoba and North America
- 2) Learn how early detection can lead to improved outcomes and when a patient is high risk
- 3) Learn how to appropriately screen patients and when to refer to nephrology

### Case

- 45 yo Male, PMH of newly diagnosed diabetes with recent A1C 9.5%, obesity, and is a smoker. His labs show a GFR 30, Urine ACR 300.
- 80 yo Male, PMH of DM2 for 20 years with good control, obesity, and is a smoker. His labs show GFR 30, ACR 10.
  - Do they have CKD?
  - Are they at high risk for ESRD?
  - Should they see a nephrologist?

### **DEFINITIONS**

### CKD:

Abnormal structure or function

- >3 months
- •Proteinuria, Hematuria
- Markers of kidney damage\*
- •eGFR <60 ml/min/m<sup>2</sup>

### Kidney Failure (ESRD):

Requiring kidney replacement therapy >3 months

- Facility Hemodialysis
- Home PD or HD
- Kidney Transplantation

\*nephrotic syndrome, nephritic syndrome, tubular syndromes, urinary tract symptoms, asymptomatic urinalysis abnormalities, asymptomatic radiologic abnormalities, hypertension due to kidney disease

### Case

• 45 yo Male, PMH of newly diagnosed diabetes, obesity, and is a smoker. His labs show a GFR 30, Urine ACR 300.

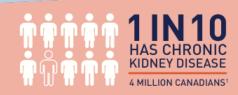
Do they have CKD?

 80 yo Male, PMH of DM2 for 20 years with good control, obesity, and is a smoker. His labs show GFR 30, ACR 10.

Do they have CKD?



WHAT YOU NEED TO KNOW



THE NUMBER OF CANADIANS LIVING WITH END-STAGE KIDNEY DISEASE HAS GROWN

**135**% SINCE 2008\*



of NEW PATIENTS
ARE UNDER THE
AGE OF 65\*





**57.5**% ARE ON DIALYSIS

42.5%
HAVE A FUNCTIONING
TRANSPLANT



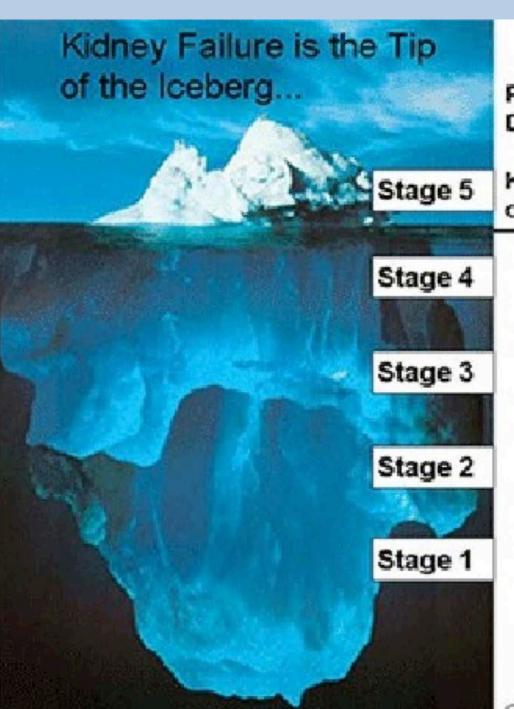
DIABETES

RENAL VASCULAR DISEASE (Including High Blood Pressure)



ASSOCIATED WITH DIALYSIS CAN AMOUNT TO 12.5% OF SOME PATIENTS' INCOME 2





Prevalence of Chronic Kidney Disease (CKD):

Kidney Failure/End-stage kidney disease (GFR <15): 400,000 2%

GFR 15-29: 300,000 1%

GFR 30-59: 7,400,000

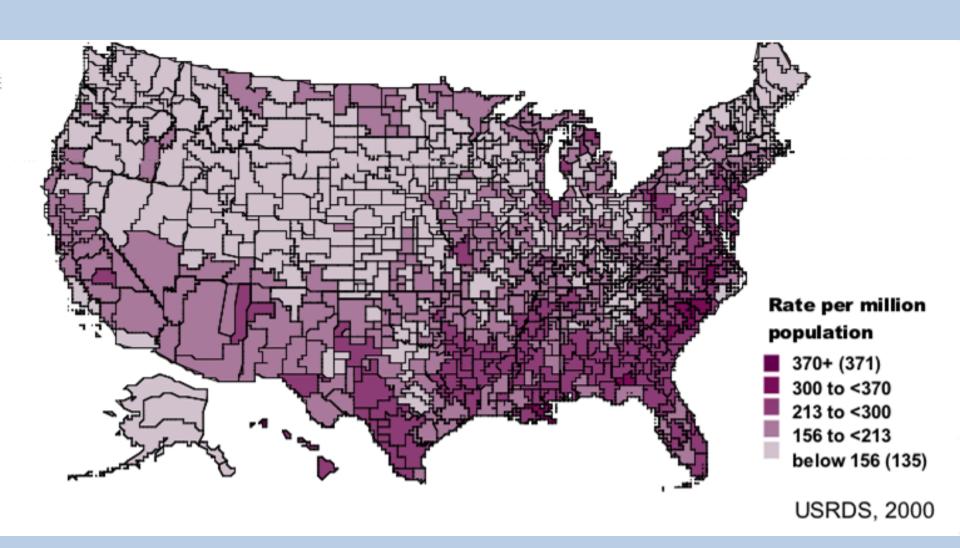
Kidney damage & GFR 60-89: 5,700,000 30%

Kidney damage & GFR >90: 5,600,000 30%

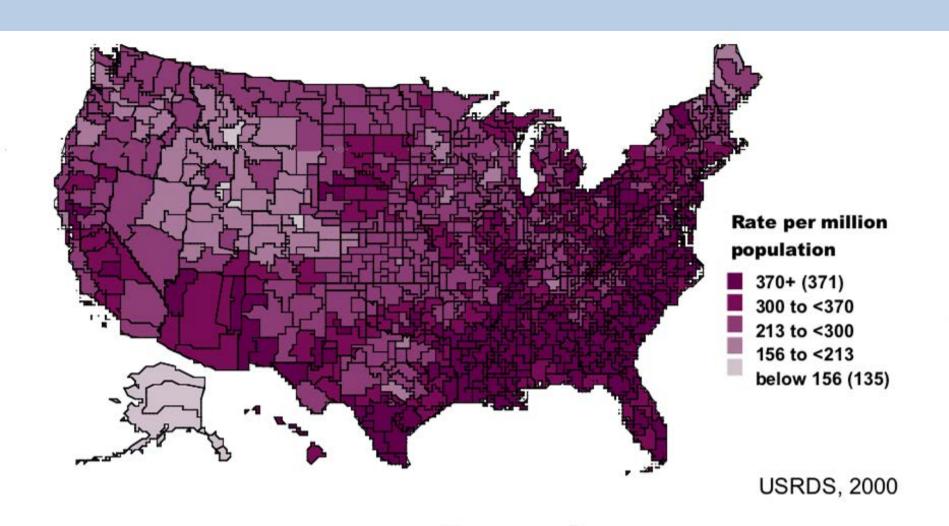
19 million Americans with CKD 8 million Americans with GFR<60

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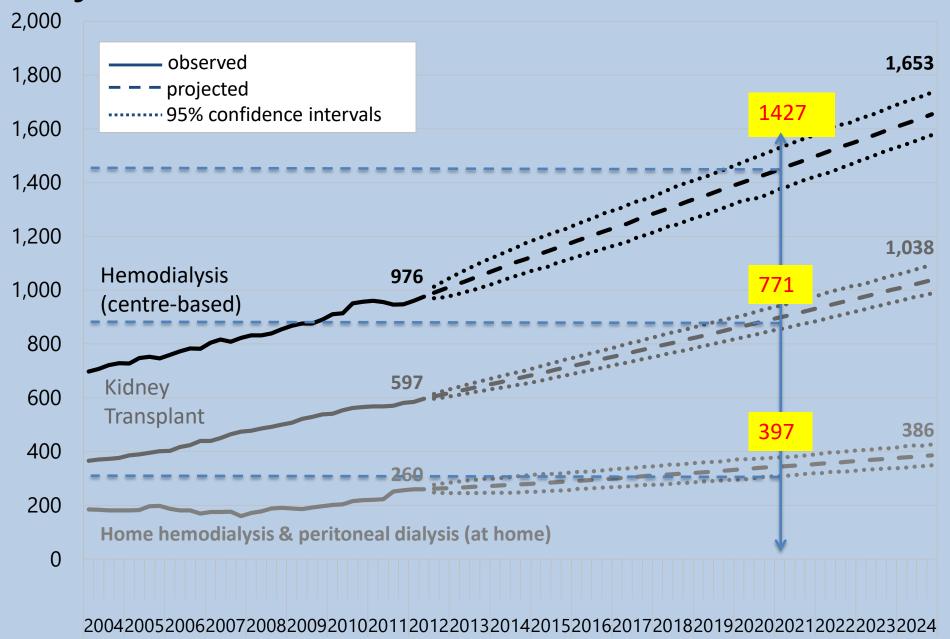
# Incidence of Kidney Failure (per million population, <u>1990</u>, from HSA)



# Incidence of Kidney Failure (per million population, **2000**, by HSA)



### **Projected Number of Manitobans with ESKD to 2024**



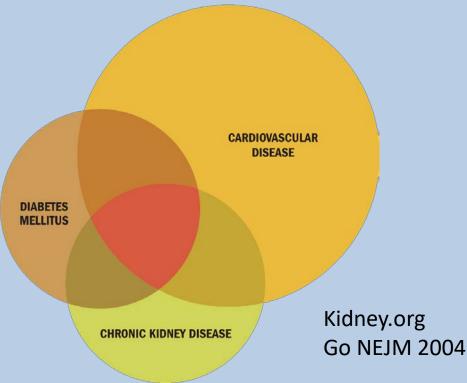
### CKD as a Public Health Issue

- Mortality, CV Events, Hospitalizations strongly related to eGFR
- 6 fold increase in mortality rate with DM + CKD (ie. multipliers)
- Disproportionately affects African Americans and Hispanics and Indigenous Peoples

Table 2. Adjusted Hazard Ratio for Death from Any Cause, Cardiovascular Events, and Hospitalization among 1,120,295 Ambulatory Adults, According to the Estimated GFR.\*

Estimated GFR	Any Death from Cardiovascular Any Any Cause Event Hospitalization adjusted hazard ratio (95 percent confidence interval)		
≥60 ml/min/1.73 m²†	1.00	1.00	1.00
45-59 ml/min/1.73 m <sup>2</sup>	1.2 (1.1-1.2)	1.4 (1.4–1.5)	1.1 (1.1–1.1)
30–44 ml/min/1.73 m²	1.8 (1.7-1.9)	2.0 (1.9–2.1)	1.5 (1.5-1.5)
15-29 ml/min/1.73 m <sup>2</sup>	3.2 (3.1-3.4)	2.8 (2.6-2.9)	2.1 (2.0-2.2)
<15 ml/min/1.73 m²	5.9 (5.4–6.5)	3.4 (3.1-3.8)	3.1 (3.0-3.3)

<sup>\*</sup> The analyses were adjusted for age, sex, income, education, use or nonuse of dialysis, and the presence or absence of prior coronary heart disease, prior chronic heart failure, prior ischemic stroke or transient ischemic attack, prior peripheral arterial disease, diabetes mellitus, hypertension, dyslipidemia, cancer, a serum albumin level of 3.5 g per deciliter or less, dementia, cirrhosis or chronic liver disease, chronic lung disease, documented proteinuria, and prior hospitalizations.

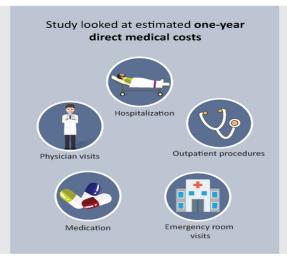


<sup>†</sup> This group served as the reference group.

### What does it cost to care for someone with kidney disease in Canada?

Alberta administrative health data from 219,641 adults\* with kidney disease not on dialysis or transplanted





Health care costs for patients with kidney disease are more than twice that of other Canadians



Cost of care was higher for people with more co-morbidities and more severe kidney disease

The cost of care for Canadians with kidney disease (not on dialysis) is high, estimated to be \$32 billion per year Further investment in effective programs and treatments to slow progression of kidney disease is recommended



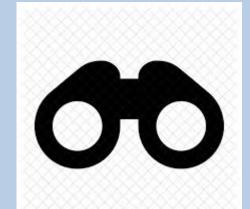


Manns et al. 2019, CJKHD

### 2019 - \$264 billion spent (12% budget)

Steps	Cost
Identification of high risk and early disease	\$
Prevention of progression	\$\$
Transplant	\$\$\$
Dialysis	\$\$\$\$

# The CURRENT CKD Journey







### FIND

Primary care screening

### DECIDE

- Risk of progression
- Refer or not refer
- When to prepare for dialysis/transplant
- When to start dialysis

# **TREAT**

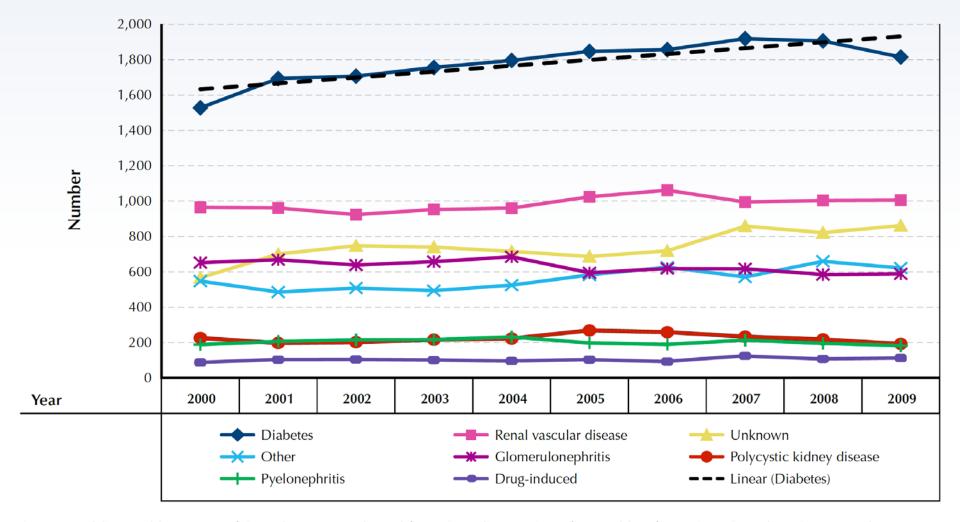
- Blood pressure
- Glucose management
- Multidisciplinary Care
- Dialysis/Transplant

# **Risk Factors**

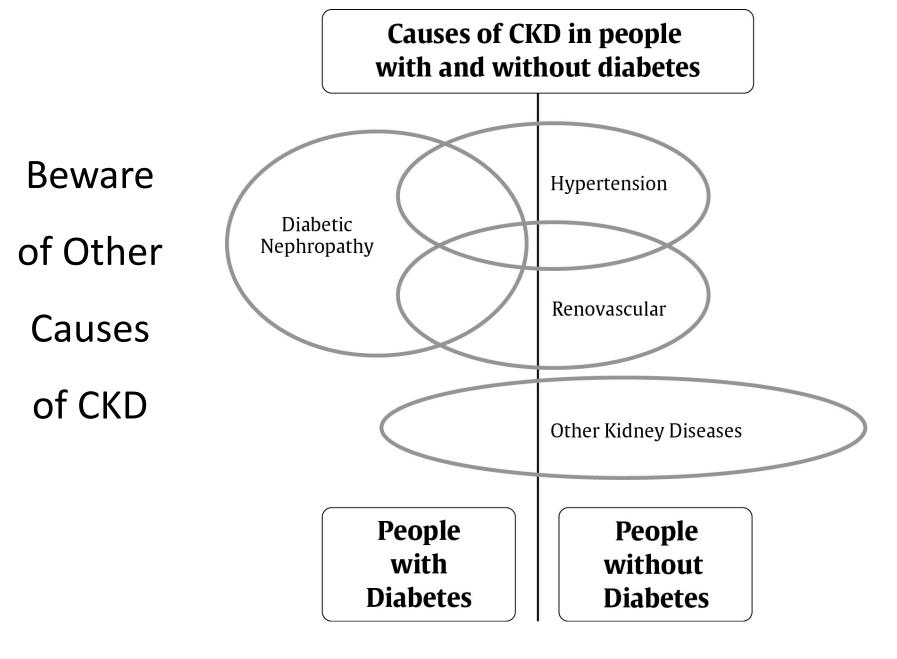
Modifiable	Non Modifiable	
Diabetes	Family History	
Hypertension	Age > 60	
AKI	Race	
NSAIDs		
CV Disease		

### Diabetes is #1 Cause of New Cases of ESRD

Figure 2-3. Number of incident cases of end-stage renal disease, by primary diagnosis, Canada, 2000 to 2009



Source: Public Health Agency of Canada (2011); adapted from Canadian Institute for Health Information. Canadian Organ Replacement Register Annual Report: Treatment of End-Stage Organ Failure in Canada, 2000 to 2009. 2011. Ottawa.

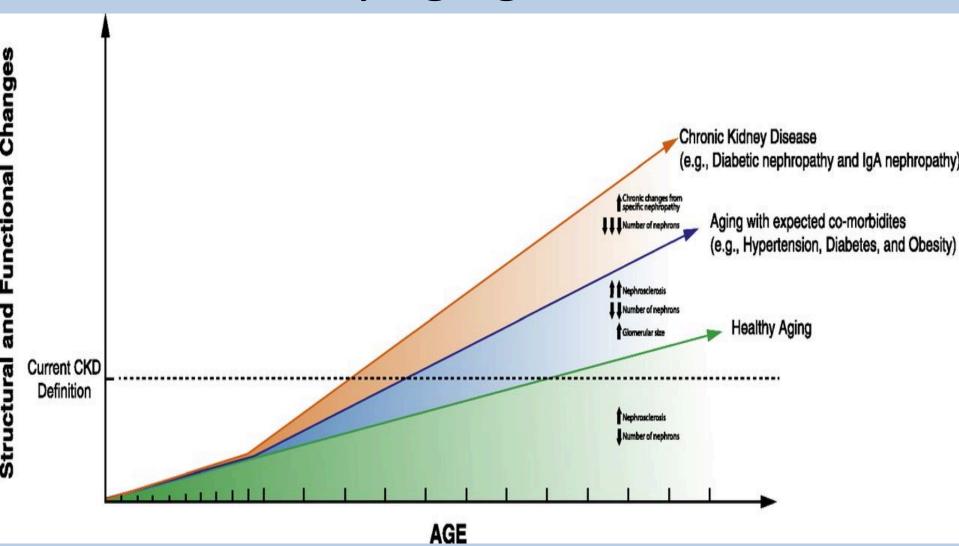


# Screening

$$f(\omega): \int_{\mathbb{R}^{2}} f(x) e^{-4\pi x x \omega} dx \xrightarrow{\mathcal{A}_{0}} (1 \xrightarrow{\mathcal{A}_{0}} \psi = H \psi)$$

$$= \left(\frac{2\pi}{2} + v \cdot \nabla v\right) \cdot \nabla p \cdot \nabla T \cdot \ell + \sum_{n=1}^{\infty} \frac{2\pi}{2} \cdot \left(\frac{\pi}{2}\right) \cdot \left(\frac{\pi}{2}$$

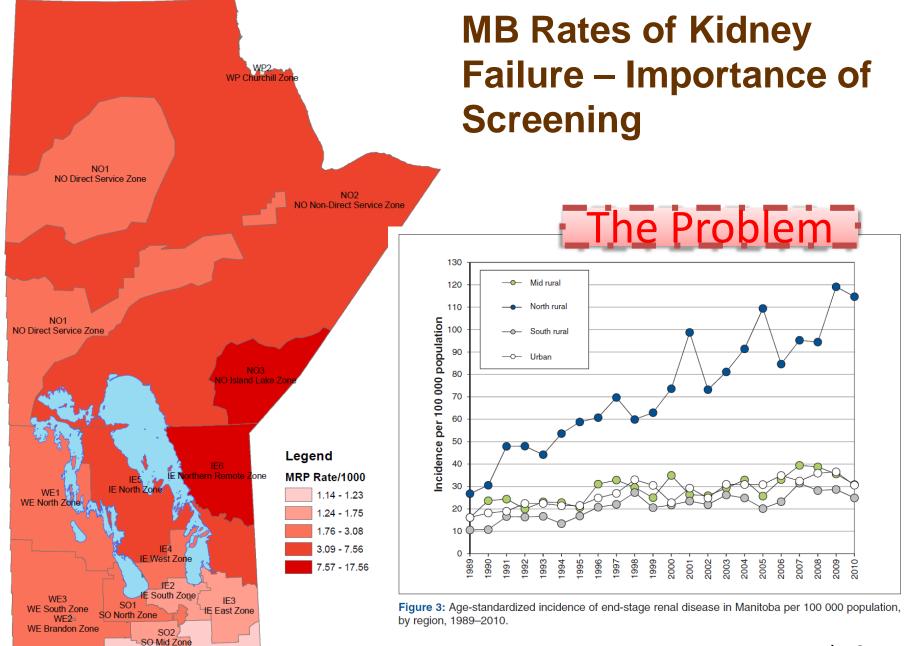
# Healthy aging and eGFR



Hommos JASN 2017

# Gaps in CKD Screening

					1
Indicat	tor	Total	Events	%	CIFª (%)
Screen	ning/recognition of CKD		8 1 5 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1		
I	% of patients with an initial eGFR <60 mL/min/1.73 m <sup>2</sup> who received a repeat SCr test in the following 6 months	218 309	107 483	49	50
2	% of patients with an initial eGFR <60 mL/min/1.73 m <sup>2</sup> who received a urine albumin-to-creatinine test in the following 6 months (including the day of the initial eGFR)	218 309	120 876	55	65
3	% of patients with an initial ACR >3 mg/mmol who received a repeat urine albumin-to-creatinine test in the following 6 months	131 178	55 583	42	42



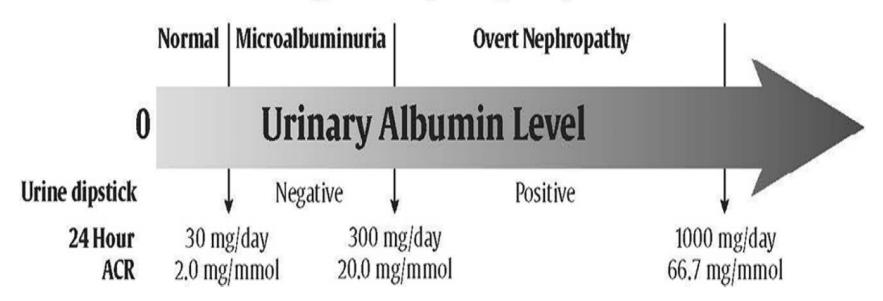
SO4

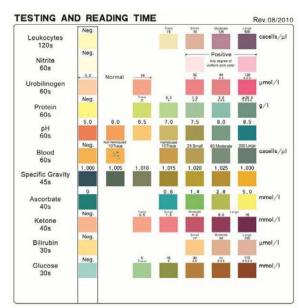
SO East Zone

SO3

SO West Zone

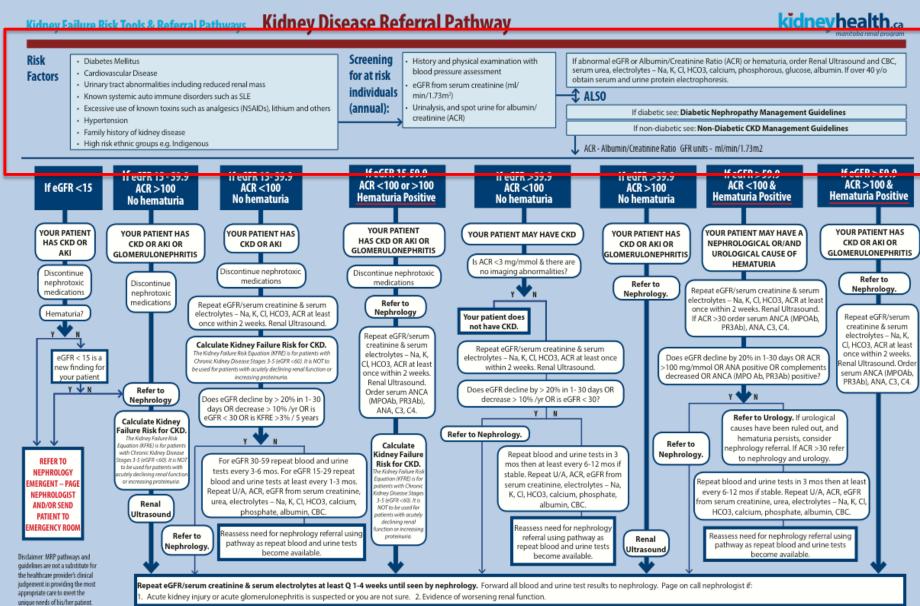
## Stage of Nephropathy





Guidelines.diabetes.ca

# Screening – who, when, how often



### **Risk Factors**

Diabetes
CV Disease
Urinary Tract Abnormalities
Reduced Renal Mass
Al disease (ex. SLE)
NSAIDS, Lithium Use
Hypertension
Family History of kidney disease
High Risk Ethnic Groups ex.
Indigenous

Annual screening for at risk Individuals

Hx and Px (with BP)
eGFR
U/A and Urine ACR

If AbN order a renal U/S, CBC, serum Ur, Na, K, Cl, HCO3, Ca, PO4, Glucose, Alb. If age>40 get SPEP and SFLC

If Diabetes, see Diabetic Nephropathy Management Guidelines. If Non Diabetic see Non-Diabetic CKD Management Guidelines

#### Kidney Failure Risk Tools & Referral Pathways

#### kidneyhealth.ca manitoba renal program

### **Diabetic Nephropathy Management Guidelines**

**Use concomitantly with Kidney Disease Referral Pathway** If ACR Start ACE-I or ARB (unless Screen 2 mg/mmol (on 2/3 eGFR < 30 or BP <110/70) annually N specimens) Repeat ACR in 4-6 weeks Is ACR Increase dose of < 2 mg/mmol or ACE-I or ARB reduced ≥ 50% ĮΥ Check BP Q 3 mos. Repeat blood and urine BP < 130/80? tests at intervals per Kidney Disease Referral Pathway Increase dose of ACE-I/ARB and/or add thiazide diuretic BP<130/80 4-6 weeks later? Add CCB and titrate to max tolerated dose and if required add other meds such as B-Blocker or alpha blockers to achieve target. Consider consult to Hypertension clinic or Nephrology if target not achieved

#### Interventions and Targets for Diabetic Nephropathy:

- · Regular exercise program
- Weight loss if obesity
- · Cessation of smoking
- Low Sodium Diet (Advise check food labels)
- Avoid NSAIDS & other nephrotoxins
- Target BP < 130/80</li>
- Target HgbA1C < 7% (check Q 3 mos)</li>

- · Consider statin treatment
- Assess medications for potential contraindication in CKD
- Adjust medication dosing for level of eGFR

when patient is on 4 medications or

on 3 but intolerant of others.

- Discontinue metformin for eGFR <30. Use with caution or discontinue for eGFR <60.</li>
- Monitor glucose closely and consider stopping long acting sulfanylurea (eg glyburide) in patients with declining eGFR.

#### Guidelines for using ACE-I and ARB:

- ACE-I/ARB are absolutely contraindicated in pregnancy. Any pre-menopausal woman on and ACE-I/ARB should be on an appropriate contraceptive method.
- Always check eGFR and serum K prior to and in 1-2 weeks of initiation or increasing dose of ACE-I/ARB.
- Expect up to a 15% decrease in eGFR. If eGFR decreases repeat again in 1-2 weeks and If eGFR decrease is stable continue ACE-I/ARB; if not, repeat eGFR again in 1-2 weeks. If eGFR continues to decrease, STOP ACE-I/ARB.
- IF serum K > 5 advise dietary K restriction.
- IF serum K > 6 advise dietary K restriction +/- prescribe diuretic if tolerated, +/prescribe K resin binder. If unsuccessful in lowering serum K to < 5.5, DECREASE OR STOP ACE-I/ARB,
- HOLD ACE-I/ARB if patient has severe vomiting/ diarrhea, or volume depletion

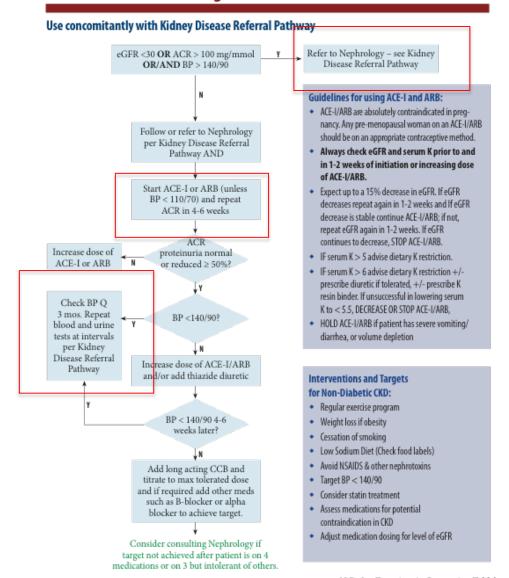
#### N.B. See 'Proteinuria Conversion Table'

Disclaimer: MRP pathways and guidelines are not a substitute for the healthcare provider's clinical judgement in providing the most appropriate care to meet the unique needs of his/her patient.

#### **Kidney Failure Risk Tools & Referral Pathways**

### kidneyhealth.ca

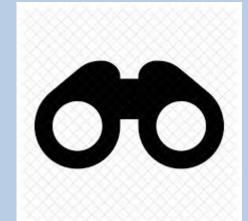
### **Non-Diabetic CKD Management Guidelines**



Disclaimer: MRP pathways and guidelines are not a substitute for the healthcare provider's clinical judgement in providing the most appropriate care to meet the unique needs of his/her patient.

N.B. See 'Proteinuria Conversion Table'

# The CURRENT CKD Journey







## **FIND**

Primary care screening

### **DECIDE**

- Risk of progression
- Refer or not refer
- When to prepare for dialysis/transplant
- When to start dialysis

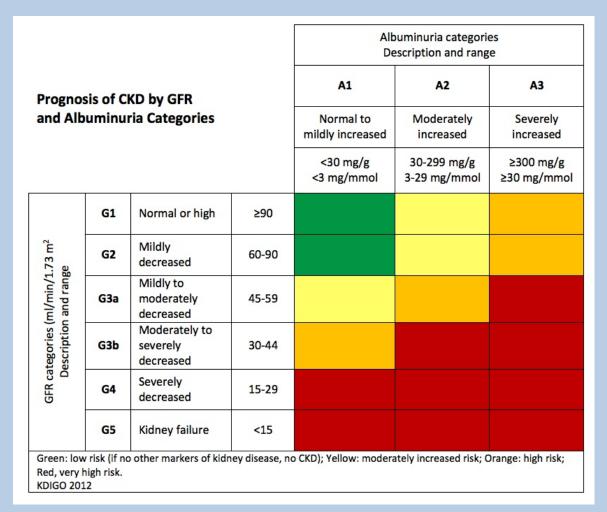
# **TREAT**

- Blood pressure
- Glucose management
- Multidisciplinary Care
- Dialysis/Transplant

# **Estimating Risk**



# Heat Map of Classification of CKD



Kidney Disease: Improving Global Outcomes (KDIGO) CKD Work Group. *Kidney Int Suppls*. 2013;3:1-150.



#### **Original Investigation**

#### The Prevalence of CKD in Rural Canadian Indigenous Peoples: Results From the First Nations Community Based Screening to Improve Kidney Health and Prevent Dialysis (FINISHED) Screen, Triage, and Treat Program

Paul Komenda, MD, MHA,<sup>1</sup> Barry Lavallee, MD,<sup>1,2</sup> Thomas W. Ferguson, MSc,<sup>1</sup> Navdeep Tangri, MD,<sup>1</sup> Caroline Chartrand, RN,<sup>2</sup> Lorraine McLeod, RN,<sup>2</sup> Audrey Gordon, MBA,<sup>3</sup> Allison Dart, MD,<sup>4</sup> and Claudio Rigatto, MD<sup>1</sup>

			Albuminuria (mg/g)			
			A	\1	A2	A3
			Optimal to high-normal		Optimal to high-normal High	
			<10	10-<30	30-<300	≥ 300
	G1a	≥105	26.4% (24.0%-28.7%)	18.6% (16.6%-20.7%)	10% (8.4%-11.6%)	1.9% (1.1%-2.6%)
	G1b	90- 104	10.2% (8.6%-11.8%)	8.2% (6.8%-9.7%)	4.8% (3.6%-5.9%)	0.7% (0.2%-1.1%)
	G2a	75-89	5.3% (4.1%-6.6%)	3.2% (2.3%-4.1%)	2.6% (1.8%-3.5%)	1% (0.5%-1.6%)
eGFR (mL/	G2b	60-74	1.6% (0.9%-2.2%)	1.0% (0.4%-1.5%)	0.9% (0.4%-1.4%)	0.3% (0%-0.6%)
min/ 1.73 m <sup>2</sup> )	G3a	45-59	0.4% (0.1%-0.8%)	0.8% (0.3% <b>-</b> 1.3%)	0.3% (0%-0.6%)	0.2% (0%-0.5%)
	G3b	30-44	0.1% (0%-0.2%)	0.2% (0%-0.5%)	0.4% (0%-0.7%)	0.4% (0.1%-0.8%)
	G4	15-29	0% (0%-0%)	0% (0%-0%)	0% (0%-0%)	0.3% (0%-0.6%)
	G5	<15	0% (0%-0%)	0% (0%-0%)	0% (0%-0%)	0.1% (0%-0.4%)

Totals: (72	74.5%	19.5%	4.5%	1.5%
	.2%-76.8%)	(17.4%-21.7%)	(3.4%-5.6%)	(0.8%-2.1%)



### THE PROJECTED RISK OF KIDNEY FAILURE



### Case

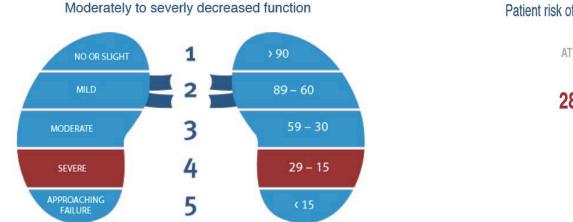
45 yo Male, PMH of newly diagnosed diabetes, obesity, and is a smoker. His labs show a GFR 30, Urine ACR 300.

Are they at high risk?

80 yo Male, PMH of DM2 for 20 years with good control, obesity, and is a smoker. His labs show GFR 30, ACR 10.

Are they at high risk?

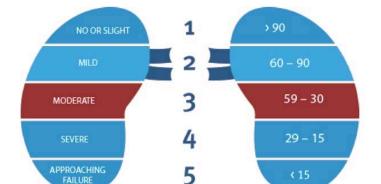
# 45 yo Male, PMH of newly diagnosed diabetes, obesity, and is a smoker. His labs show a GFR 30, Urine ACR 300.



Patient risk of progression to kidney failure requiring dialysis or transplant:



80 yo Male, PMH of DM2 for 20 years with good control, obesity, and is a smoker. His labs show GFR 30, ACR 10.



Mildly to moderately decreased function

Patient risk of progression to kidney failure requiring dialysis or transplant:



### **Kidney Failure Risk Tools & Referral Pathways**

### **Outpatient Nephrology Referral Form**



teferring MD:	Ph	Fax
atient Name:		Phone:
PHIN:A	.ddress:	
lease complete and append results of:  Past medical history  Medications list  eGFR (also serum urea, creatinine) at least two values  CBC  Urinalysis  Spot urine for ACR	EMERGENT REFERRAL (< 24 HOURS) DO NOT FAX REFERRAL	eGFR <15 mL/min with indications (see referral pathway) for emergent dialysis PAGE NEPHROLOGIST ON CALL AT: Health Sciences Centre (204) 787-2071 St. Boniface Hospital (204) 237-2053 Brandon
onsider ordering and send reports when available: Serum and Urine Protein Electrophoresis (> 40 years of age) Kidney Ultrasound	NON- EMERGENT	(204) 578-4000 or (204) 571-7139  Reason for Urgency (check all that apply)  □ eGFR <30 mL/min
REFERRAL SITE PREFERENCE (check one)  St. Boniface Hospital - Fax to (204) 233-2770  Health Sciences Centre - Fax to (204) 787-7366  Seven Oaks Hospital - Fax to (204) 697-4204  Brandon Regional Health Centre Fax to (204) 726-8797  Dr		□ KFRE >3% / 5 year     □ eGFR decline by >20% in 1-30 days (acute kidney injury)     □ Proteinuria (ACR >100 mg/mmol)     □ Hematuria, suspected GN (eGFR or proteinuria criteria below, or ANA>1:80, or decreased complements, or ANCA positive or normal urological evaluation
ignature:	Data	

1 - ACR = Albumin to Creatinine Ratio. 2 - eGFR = estimated Glomerular Filtralion Rate automatically calculated from serum Creatinine by MDRD Formula.

N.B. The noted appointment wait times are target benchmarks, but each patient will be triaged at time of referral. Appointment times may vary. Until your patient is assessed by a nephrologist, you (referring physician) are responsible for monitoring (and forwarding) your patients blood and urine tests at regular intervals as suggested in the MRP Kidney Disease Referral Pathway. You should contact the nephrologist with any new concerns.

### Case

45 yo Male, PMH of newly diagnosed diabetes, obesity, and is a smoker. His labs show a GFR 30, Urine ACR 300.

Should they see a nephrologist?

80 yo Male, PMH of DM2 for 20 years with good control, obesity, and is a smoker. His labs show GFR 30, ACR 10.

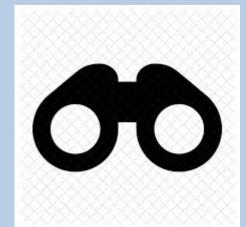
Should they see a nephrologist?

Indications for Referral to Specialist Kidney Care Services for People with CKD (KFRE is just a tool!)

eGFR < 30 ml/min
eGFR rapid decline > 10%/year or 20% in a shorter interval
K+ >6, other electrolyte abnormalities
Nephrotic Range Proteinuria +/- eGFR
Refractory Hypertension
Hematuria/Suspected GN
Structural Disease/Polycystic Kidney Disease
Pregnancy

**CALL FOR URGENT CONSULTS!!!!** 

# The CURRENT CKD Journey







## **FIND**

Primary care screening

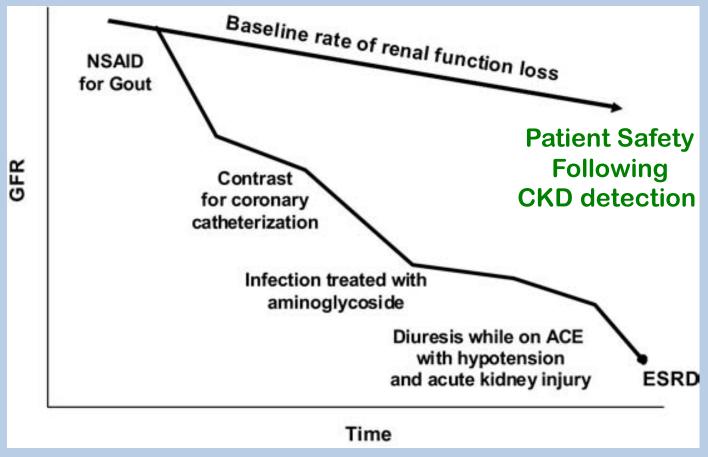
### DECIDE

- Risk of progression
- Refer or not refer
- When to prepare for dialysis/transplant
- When to start dialysis

### **TREAT**

- Blood pressure
- Glucose management
- CV Risk
- Multidisciplinary Care
- Dialysis/Transplant

# Impact of primary care CKD detection with a patient safety approach



Improved diagnosis creates opportunity for strategic preservation of kidney function

# Risk-Based Teams Approach

#### **Low Risk Patients**

### Team:

Primary care provider

### Focus:

- Lifestyle
- blood pressure

### **Medium Risk Patients**

#### Team:

- Primary Care providers
- Kidney specialist

#### Focus:

- Diagnosis
- Lifestyle
- Blood pressure
- Protein leak reduction

# Resource allocation aligned with

### **High Risk Patients**

### <u>Team:</u> Multidisciplinary Kidney Health Clinic

- Primary care provider\*
- Kidney specialist
- Kidney Nurse
- Kidney Dietitian
- Kidney Pharmacist
- Kidney Social worker

### Focus:

- Healthy Lifestyle
- Reduction of blood pressure
- Reduction of protein leak in the urine
- Treat consequences of low kidney function
- Treat related heart and diabetic diseases

### **Evidence Based Treatments for CKD**

- Sodium-Glucose **Transporter 2 Inhibitors** (SGLT-2i)
- Blood Pressure <130/80
- Renin-Angiotensin Inhibitors (ACE/ARB)
- Target optimal blood glucose (HgbA1c - 7%)
- Cardiovascular risk factor control
- **Acid-Base Control**



















The earlier we catch it, the longer time to needing dialysis

## Medication adjustments

- Declining kidney function may necessitate changes to > 30% renally-cleared drugs
- Stop NSAIDs
- Minimize polypharmacy
- Many commonly used meds need dose adjustment based on CrCl or GFR
  - Allopurinol
  - Gabapentin
  - Narcotics
  - BB
  - Digoxin
  - Antibiotics
  - MTX
  - Enoxaparin/anticoagulants
  - Diabetic meds

А	ACEi
D	Diuretics
M	Metformin
Α	ARB
N	NSAIDs
S	SGLT2 inhibitors

# Thank you!

