



XIX CONGRESSO NAZIONALE AMD

Roma, 29 maggio - 1 giugno 2013
Rome Marriott Park Hotel

La definizione e classificazione dell'IRC

A. Guarnieri

ASO S. Croce e Carle - CN
S.C. Nefrologia e Dialisi

Definizione e classificazione della malattia renale cronica

Danno renale presente da almeno tre mesi, definito da anomalie funzionali o strutturali del rene, con o senza alterazioni del filtrato glomerulare, con implicazioni per la salute.

Table 2 | Criteria for CKD (either of the following present for > 3 months)

Markers of kidney damage (one or more)	Albuminuria (AER \geq 30 mg/24 hours; ACR \geq 30 mg/g [\geq 3 mg/mmol]) Urine sediment abnormalities Electrolyte and other abnormalities due to tubular disorders Abnormalities detected by histology Structural abnormalities detected by imaging History of kidney transplantation
Decreased GFR	GFR $<$ 60 ml/min/1.73 m ² (GFR categories G3a-G5)

Abbreviations: CKD, chronic kidney disease; GFR, glomerular filtration rate.

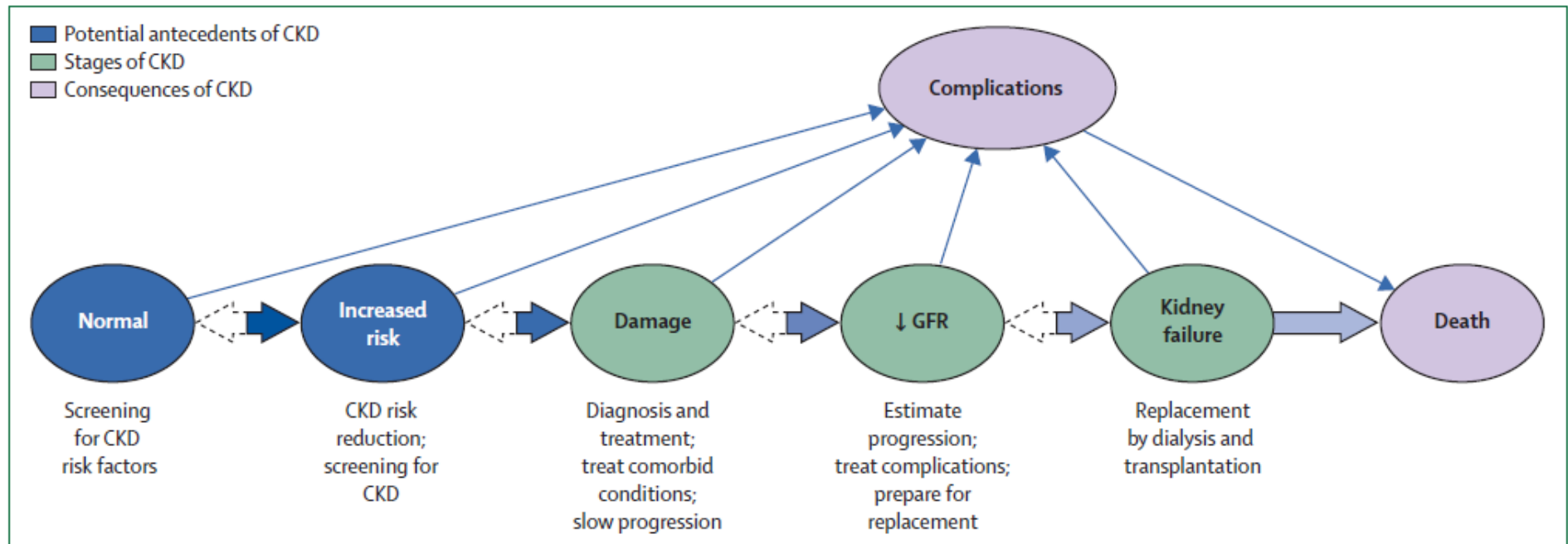
TREATMENT IN GENERAL PRACTICE

This article is one of a series on the management of some of the renal diseases met with in general practice. The first group of articles on Treatment in General Practice, thirty-five in number, was republished in book form last March by H. K. Lewis & Co., and a second volume will appear in the autumn.

CHRONIC NEPHRITIS AND URAEMIA

BY

ROBERT PLATT, M.D., F.R.C.P.



Markers di danno renale

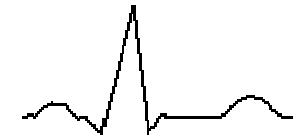
- Albuminuria



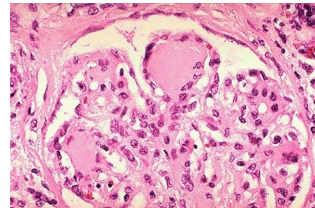
- Anomalie del sedimento urinario



- Anomalie degli elettroliti secondarie a danni tubulari



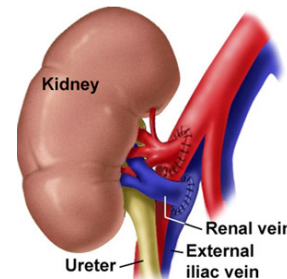
- Alterazioni istologiche



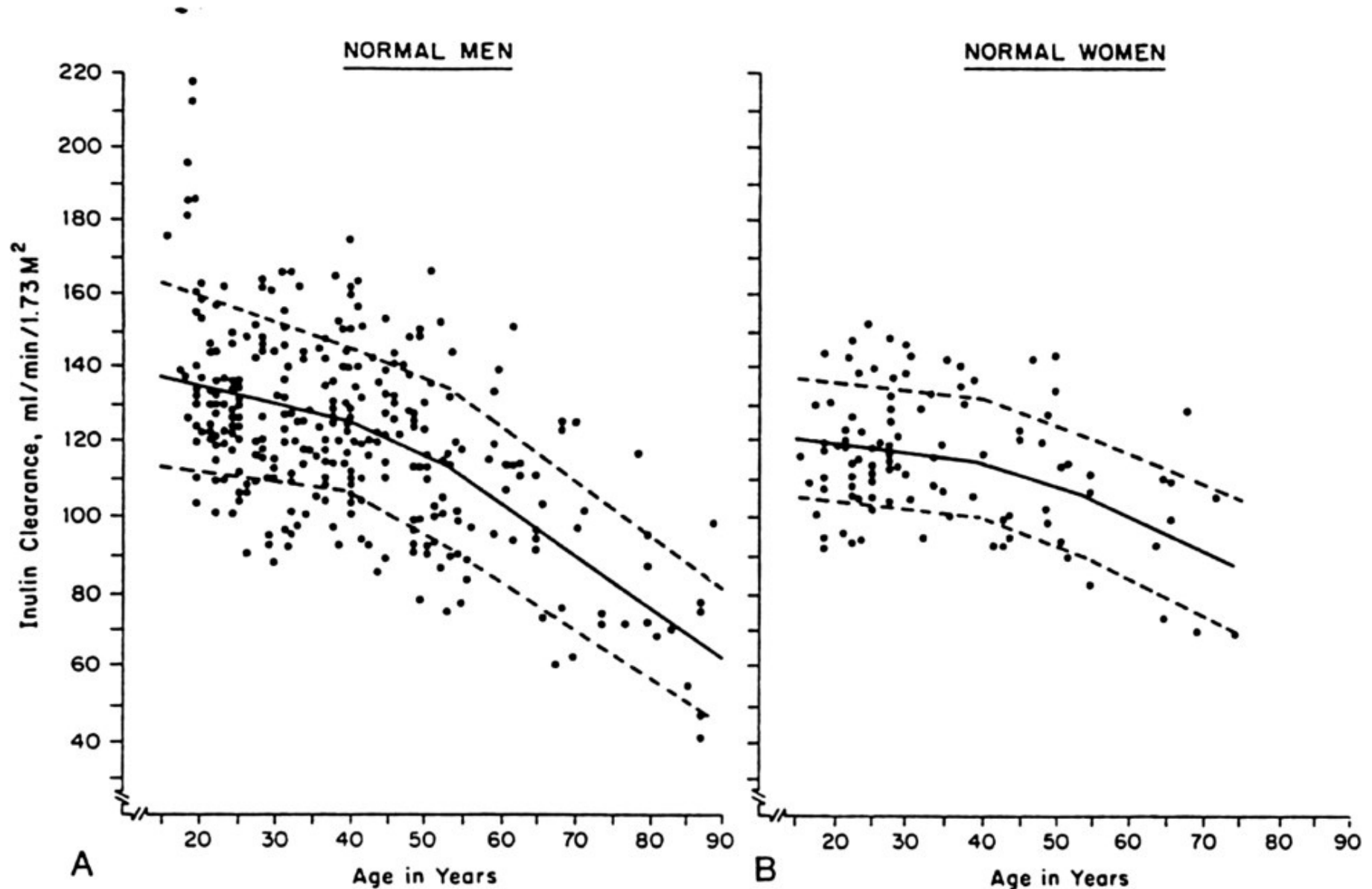
- Alterazioni evidenziabili con tecniche di immagine



- Storia di trapianto renale



IRC = GFR < 60 mL/min/1.73mq



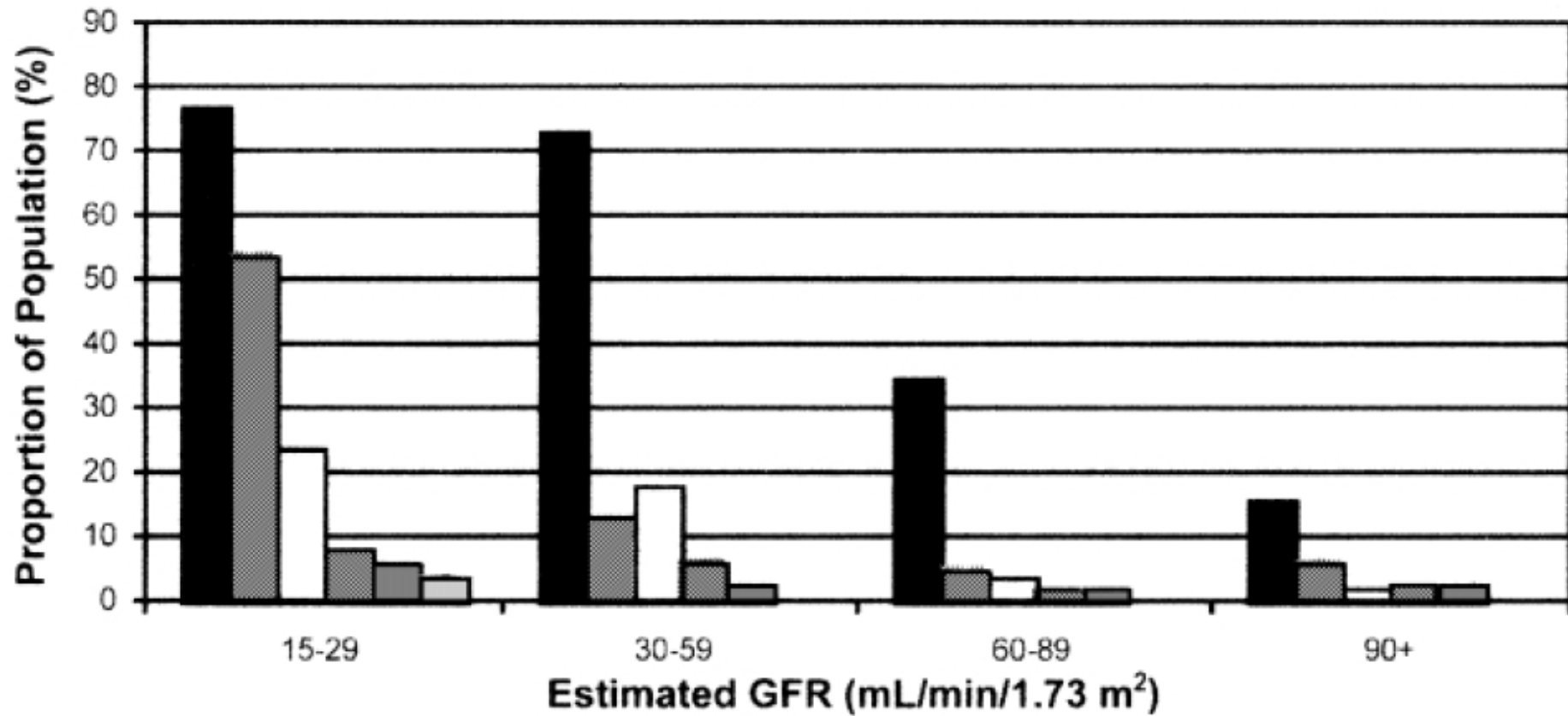
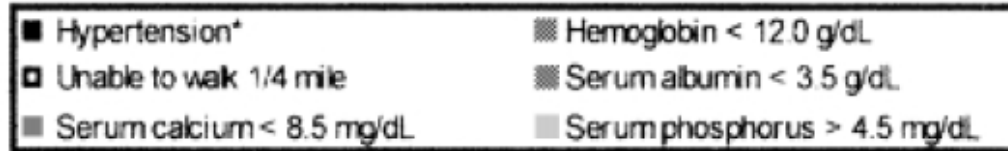
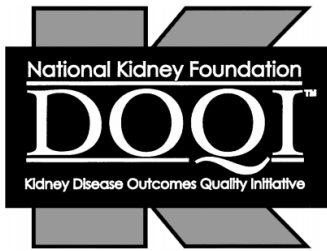
Plasma Metabolomic Profiles in Different Stages of CKD

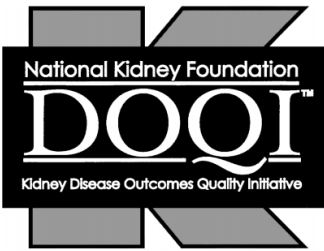
Table 2. Ratios of significant changes of specific metabolites by stage of CKD ($P \leq 0.05$), mean values, and P values

	Fold of Change			Mean \pm SD			P Value		
	Stages 3/2	Stages 4/2	Stages 4/3	CKD Stage 2	CKD Stage 3	CKD Stage 4	Stages 3/2	Stages 4/2	Stages 4/3
Altered arginine metabolism									
Dimethylarginine (SDMA + ADMA)	8.1	4.8		0.497 \pm 0.133	4.01 \pm 2.27	2.40 \pm 1.77	7.9E-05	0.01	0.17
Citrulline	1.6	1.3 ^a		0.825 \pm 0.194	1.30 \pm 0.586	1.08 \pm 0.362	0.03	0.08	
Ornithine	0.28	0.16		4.33 \pm 1.39	1.19 \pm 1.32	0.683 \pm 0.334	2.0E-4	1.4E-06	
Arginine	1.5 ^a	1.5 ^a		0.675 \pm 0.238	1.02 \pm 0.480	1.03 \pm 0.559	0.06	0.10	
Elevated coagulation/inflammation									
Fibrinopeptide A	689	827		0.002 \pm 0.001	1.03 \pm 0.765	1.24 \pm 0.795	3.0E-4	1.1E-14	
Phosphorylated fibrinopeptide A	18	45	2.5	0.045 \pm 0.017	0.818 \pm 0.683	2.06 \pm 2.62	0.002	4.4E-08	0.04
Proline-hydroxyproline	2.5	4.5		0.466 \pm 0.251	1.16 \pm 1.26	2.08 \pm 1.68	0.03	8.3E-05	
Impaired carboxylate anion transport									
γ -Glutamylleucine		1.3		0.821 \pm 0.283		1.07 \pm 0.161		0.03	
γ -Glutamylisoleucine	1.6	1.7		0.670 \pm 0.196	1.12 \pm 0.440	1.18 \pm 0.510	0.01	0.01	
γ -Glutamylglutamine	3.8	4.8		0.275 \pm 0.166	1.05 \pm 0.417	1.31 \pm 0.349	5.5E-05	8.9E-06	
γ -Glutamylphenylalanine	1.5	1.3		0.903 \pm 0.240	1.31 \pm 0.480	1.17 \pm 0.181	0.03	0.01	
CMPF	18.3 ^a	23.6		0.307 \pm 0.279	5.61 \pm 11.9	7.23 \pm 7.20	0.06	5.0E-4	
Decreased adrenal steroid hormone production									
Dehydroisoandrosterone sulfate		0.55		1.48 \pm 0.878		0.819 \pm 0.685		0.04	
4-androsten-3- β ,17- β -diol disulfate		0.26		3.91 \pm 3.76		1.02 \pm 0.958		0.01	

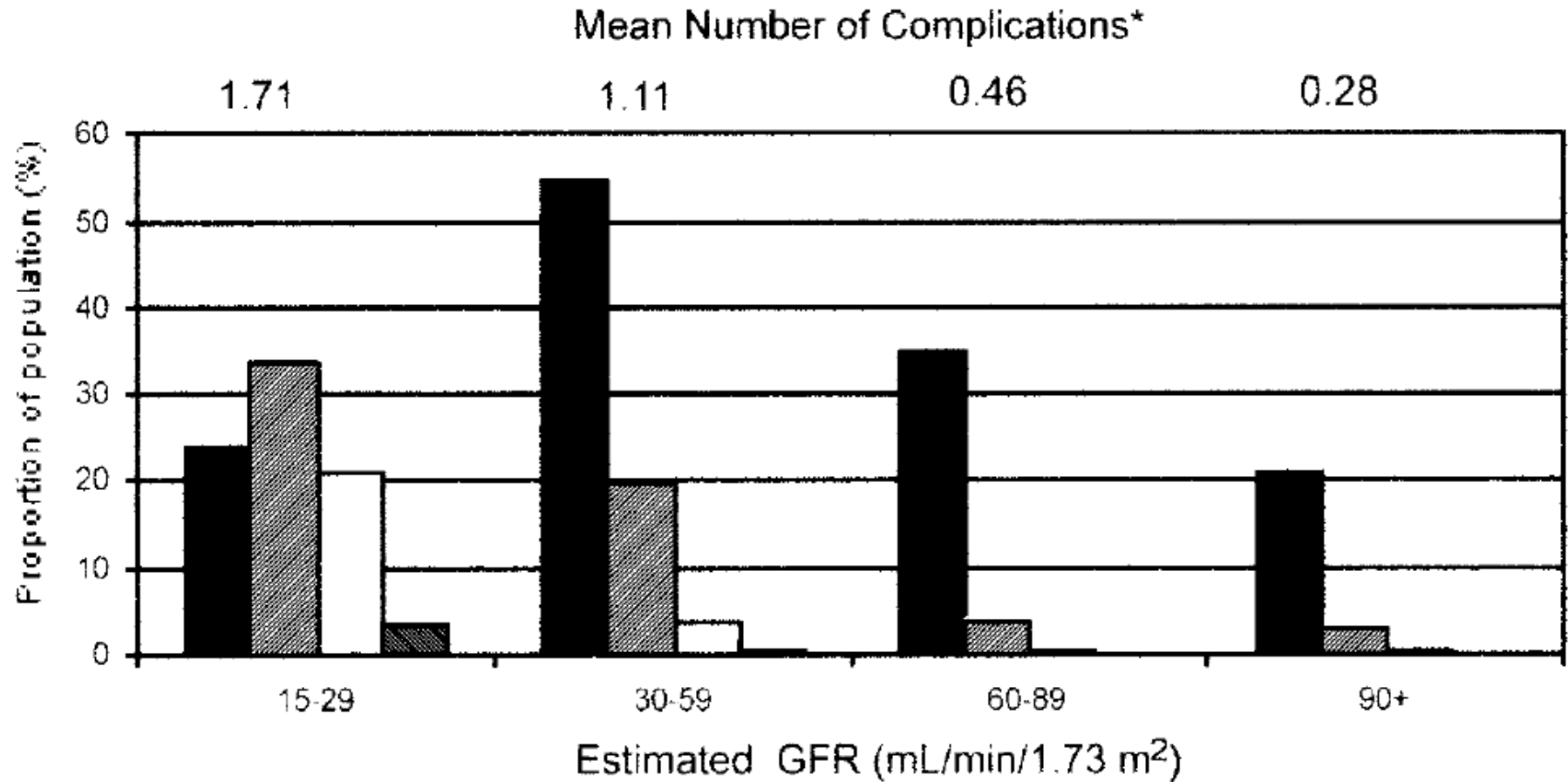
ADMA, asymmetric dimethylarginine; SDMA, symmetric dimethylarginine; CMPF, 3-Carboxy-4-methyl-5-propyl-2- furanpropanoate.

^a0.1 > P > 0.05.



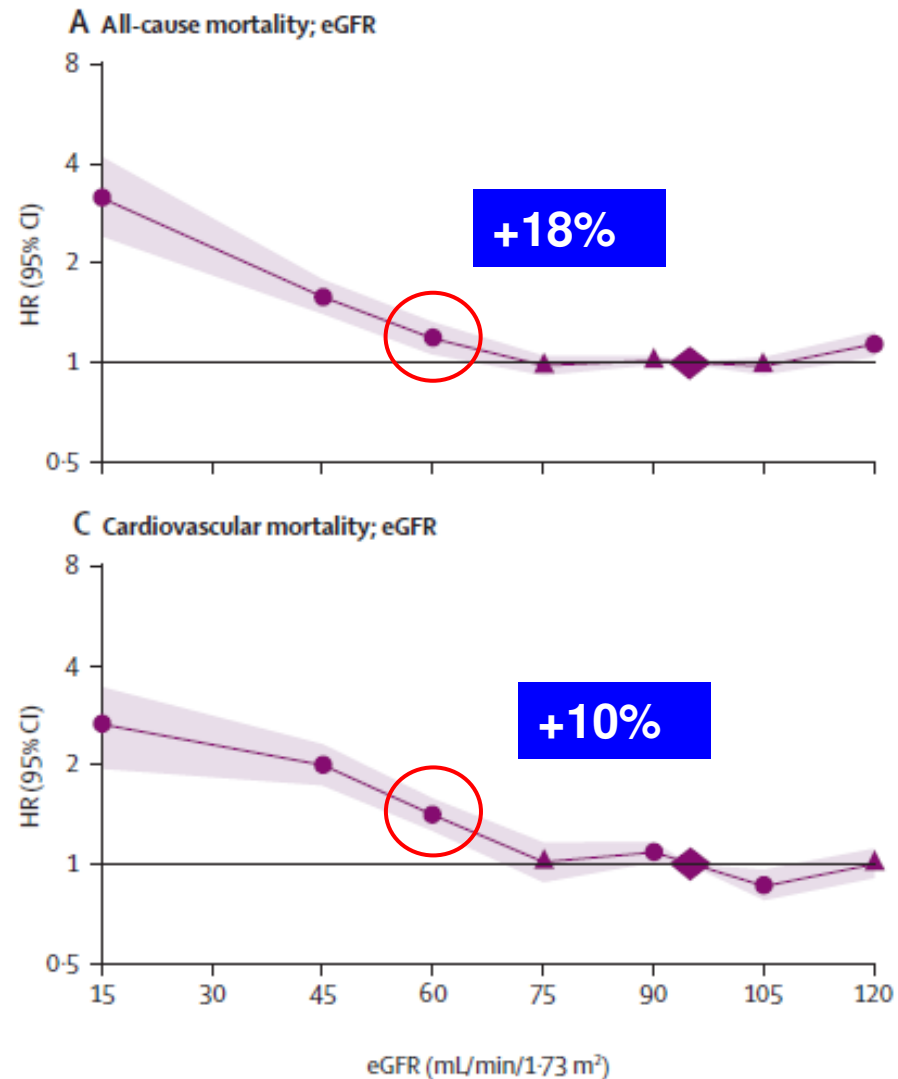


■ 1 Complication ▨ 2 Complications □ 3 Complications ▩ 4 Complications



Association of estimated glomerular filtration rate and albuminuria with all-cause and cardiovascular mortality in general population cohorts: a collaborative meta-analysis

Chronic Kidney Disease Prognosis Consortium*



Lancet 2010; 375: 2073-81



Table 1. Classification of CKD as Defined by KDOQI and Modified and Endorsed by KDIGO

Stage	Description	Classification by Severity	Classification by Treatment
1	Kidney damage with normal or \uparrow GFR	$GFR \geq 90$	} T if kidney transplant recipient
2	Kidney damage with mild \downarrow in GFR	GFR of 60-89	
3	Moderate \downarrow in GFR	GFR of 30-59	
4	Severe \downarrow in GFR	GFR of 15-29	
5	Kidney failure	$GFR < 15$ (or dialysis)	D if dialysis

CKD EPI & MDRD GFR Calculator - (With SI Units)

4 variable MDRD CKD EPI Equation (with SI Units)
using standardized serum creatinine, age, race, gender

by Stephen Z. Fadem, M.D., FACP, FASN
and Brian Rosenthal

Cockcroft-C

Male

Female

MDRD stu

GFR (ml/m

Serum creatinine

mg/dL $\mu\text{mol/L}$

1.1

GFR (ml/m

MDRD Stu

GFR (ml/m

NOTE: CKD EPI GFR is only valid with creatinine methods are traceable to IDMS

Age

30 years

Race

African American All other races*

GFR (ml/m

Gender

Male Female

CKD-EPI E

GFR (ml/m

TRACEABLE TO IDMS (What is this?) No Yes

CKD EPI Value: 104 mL/min/1.73 m² in a 30 year old African American male.

where κ is
the minimum

MDRD GFR: 95 mL/min/1.73 m² in a 30 year old African American male.

Female

Male

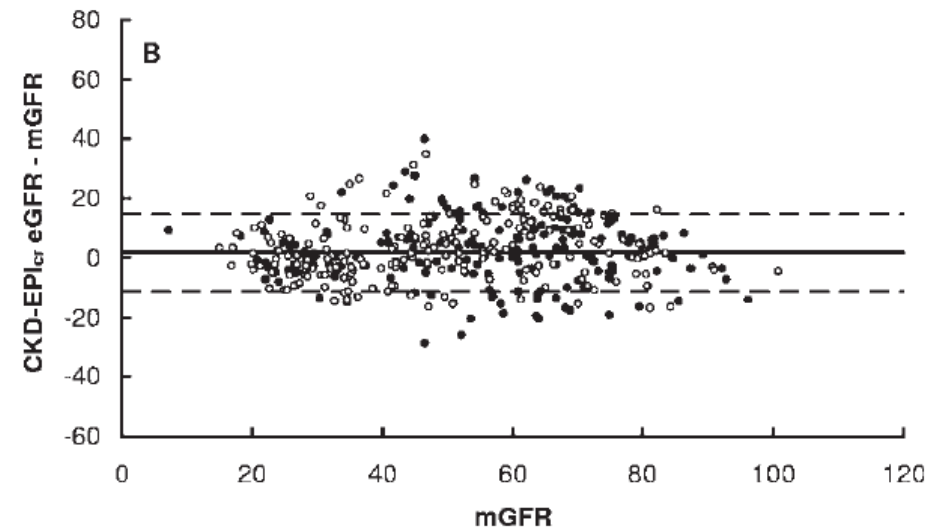
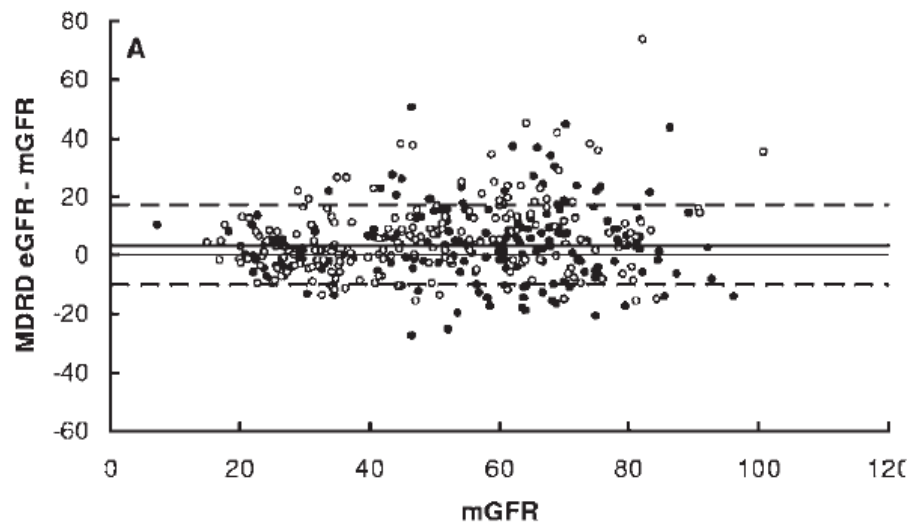
*Age in year

(Age, Race, Gender, Plasma creatinine)

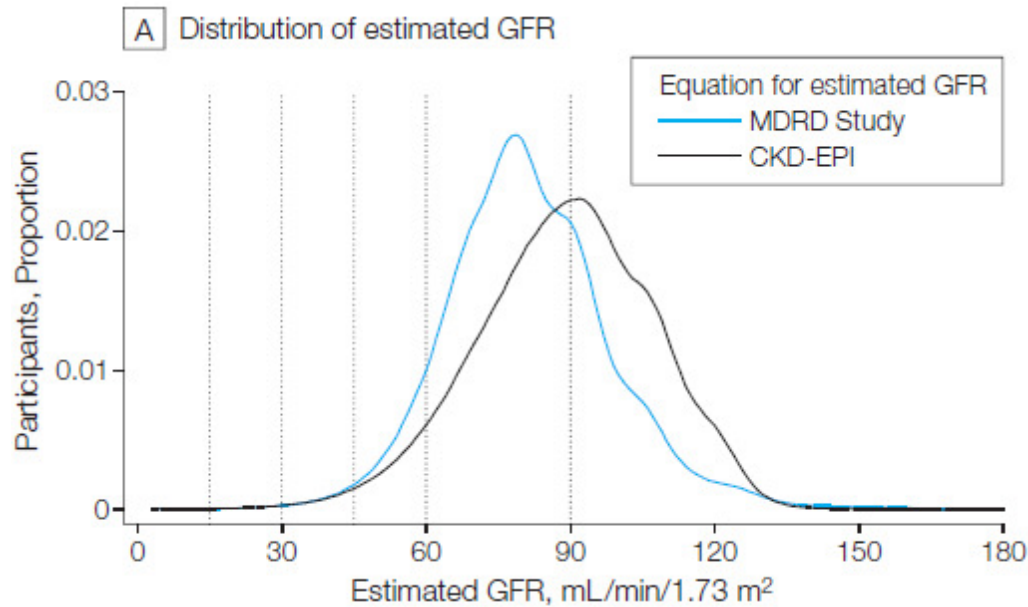
Chronic kidney disease (GFR less than 60 or kidney damage for at least three months)

Fig. 3-5. Equations for estimating glomerular filtration rate. The MDRD study equation calculator can also be found online at http://www.kidney.org/professionals/kdoqi/gfr_calculator.cfm.

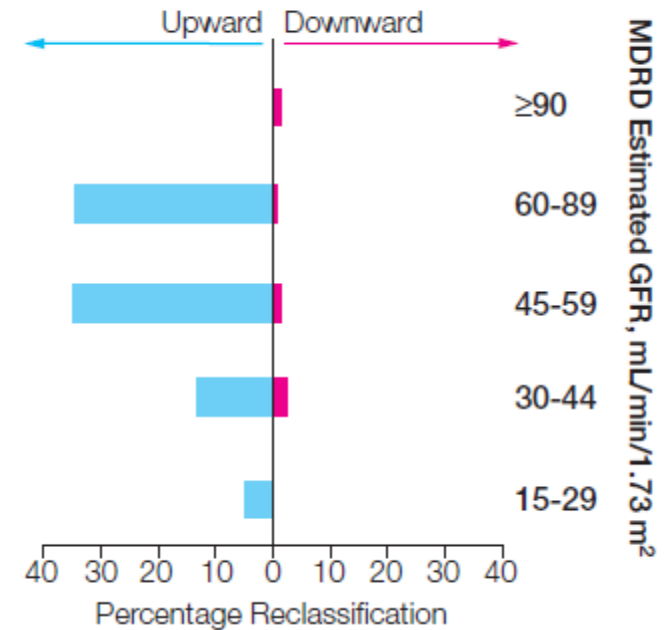
Accuracy of the MDRD (Modification of Diet in Renal Disease) Study and CKD-EPI (CKD Epidemiology Collaboration) Equations for Estimation of GFR in the Elderly



Comparison of Risk Prediction Using the CKD-EPI Equation and the MDRD Study Equation for Estimated Glomerular Filtration Rate

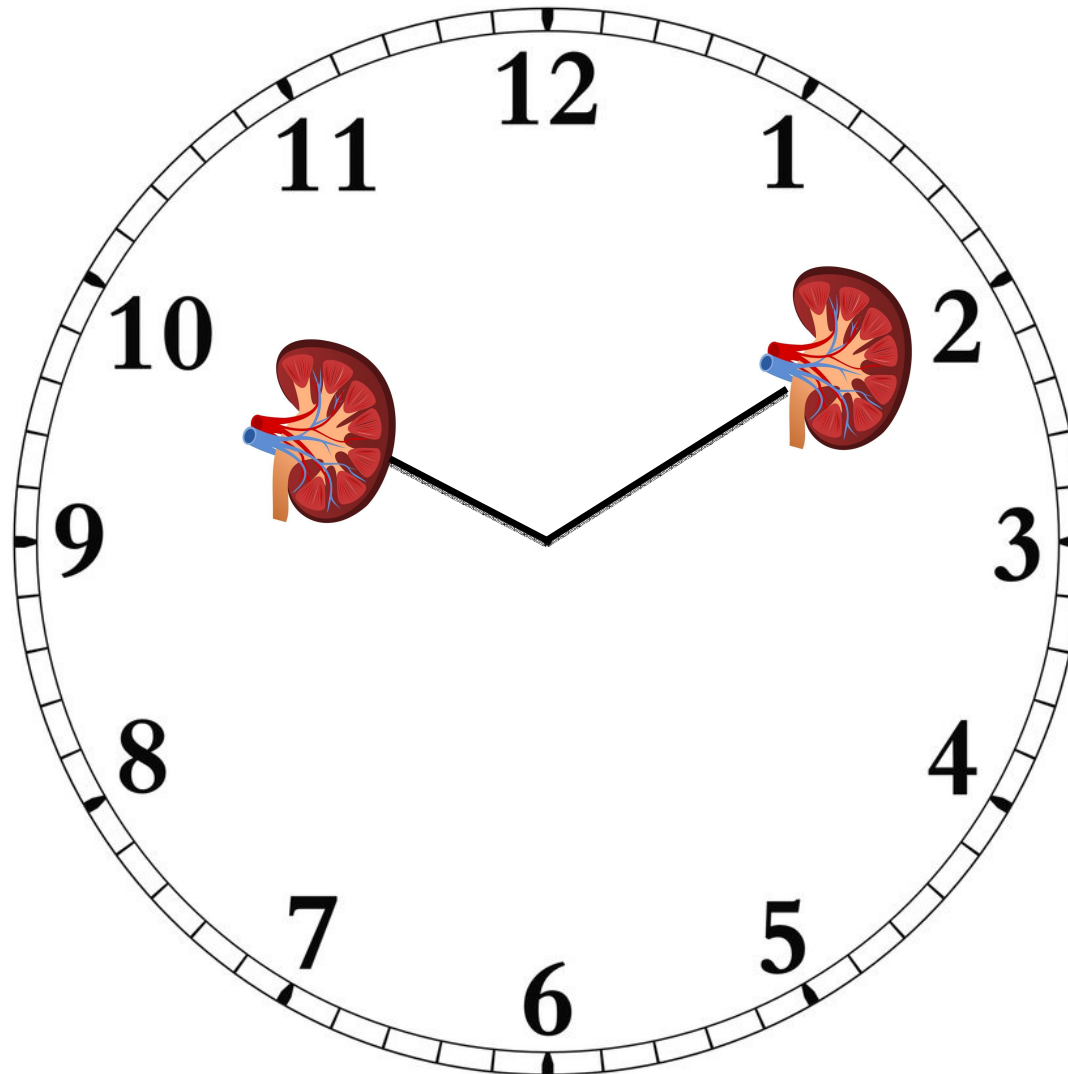


Reclassification by CKD-EPI Estimated GFR

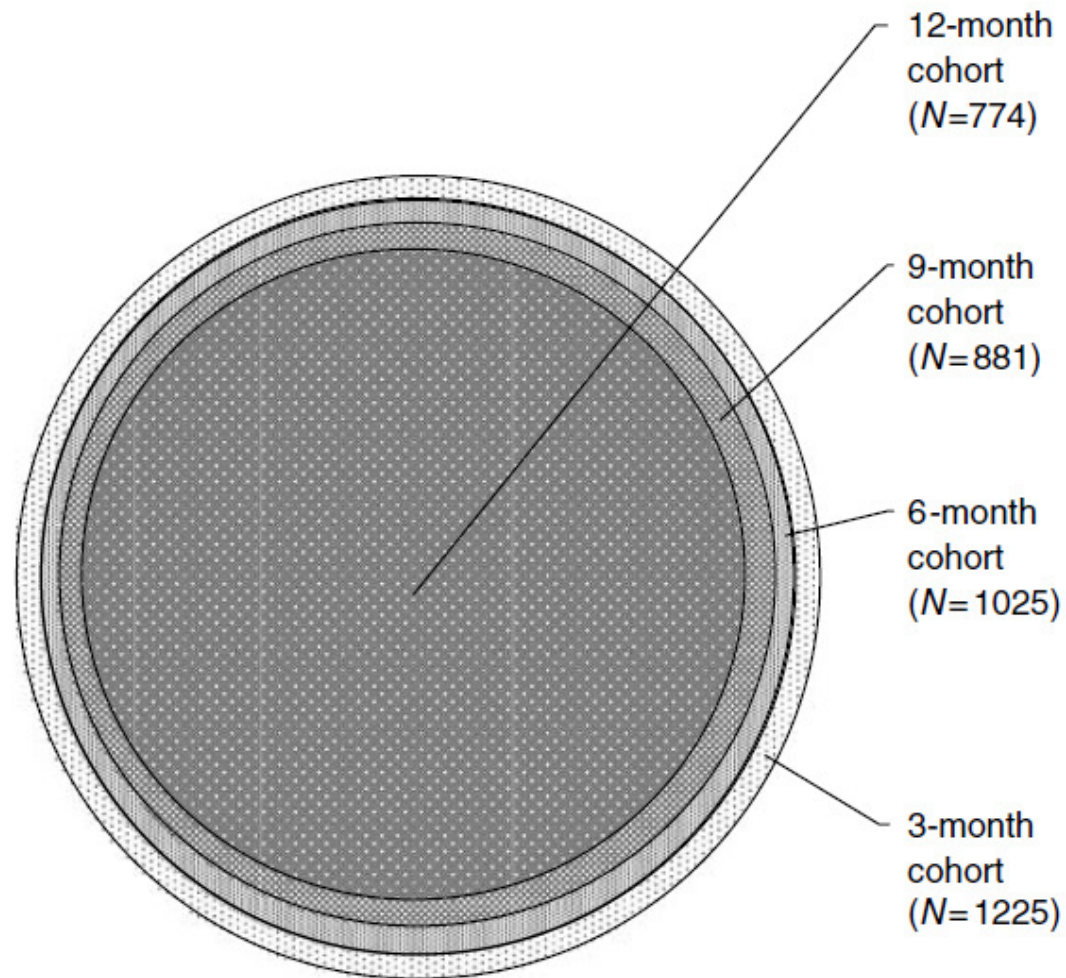


Cronicità del danno

Danno renale presente da almeno tre mesi

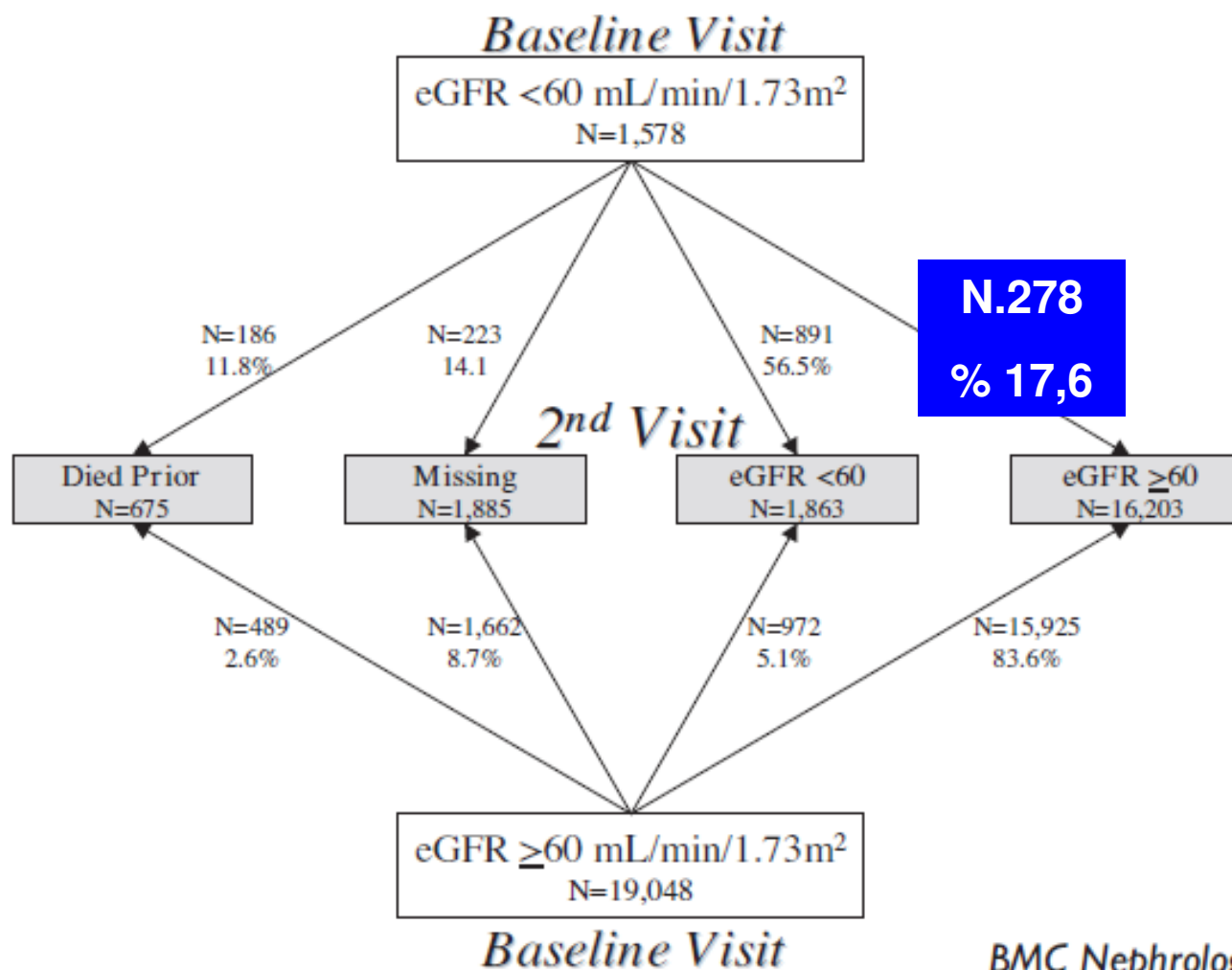


In chronic kidney disease staging the use of the chronicity criterion affects prognosis and the rate of progression

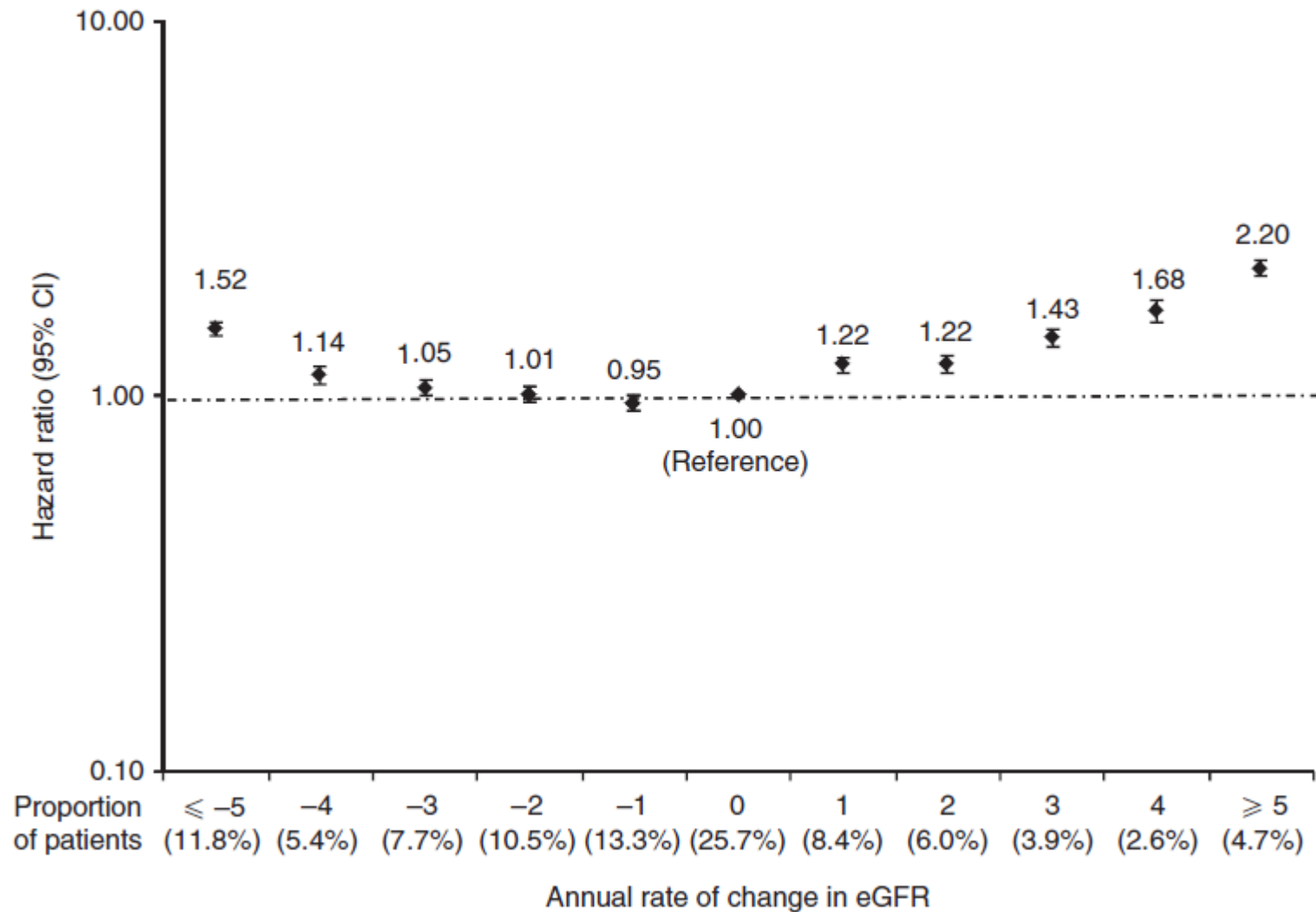


CKD classification based on estimated GFR over three years and subsequent cardiac and mortality outcomes: a cohort study

Daniel E Weiner*¹, Maria Krassilnikova², Hocine Tighiouart²,
Deeb N Salem², Andrew S Levey¹ and Mark J Sarnak¹



Change in the estimated glomerular filtration rate over time and risk of all-cause mortality

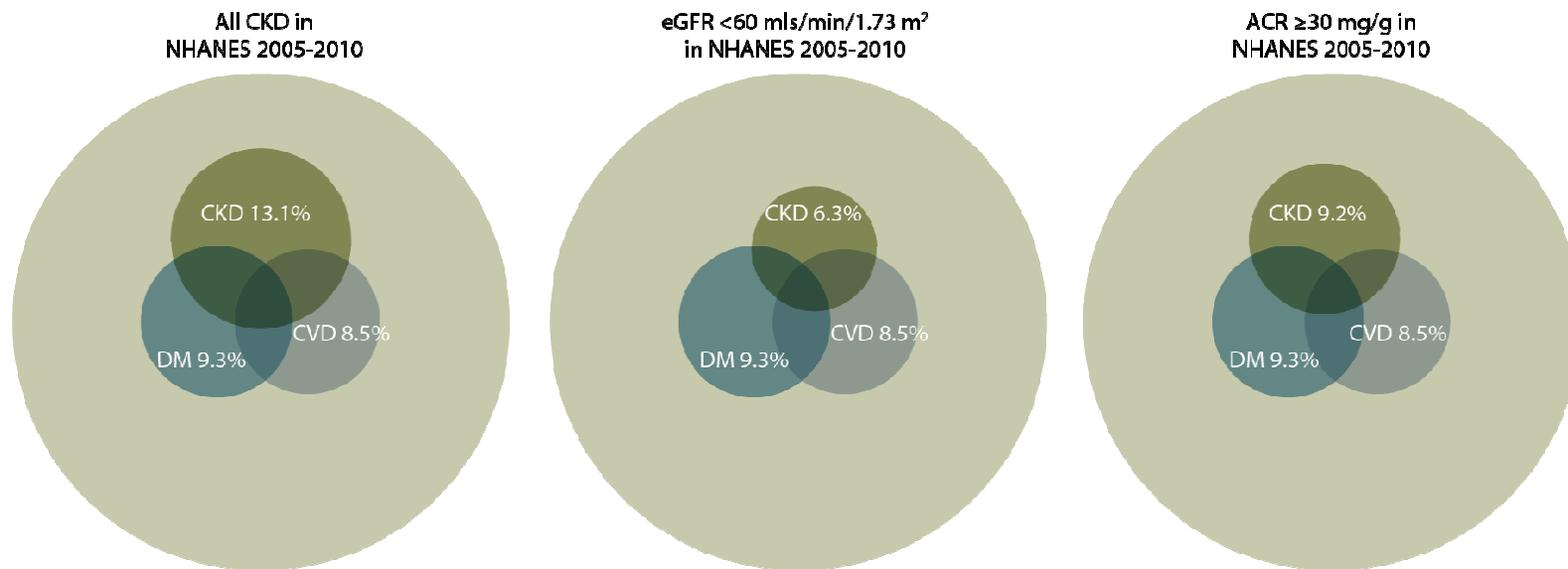


**Classe III (eGFR 30 – 59 mL/min/1.73mq):
applicabilità e caratterizzazione**



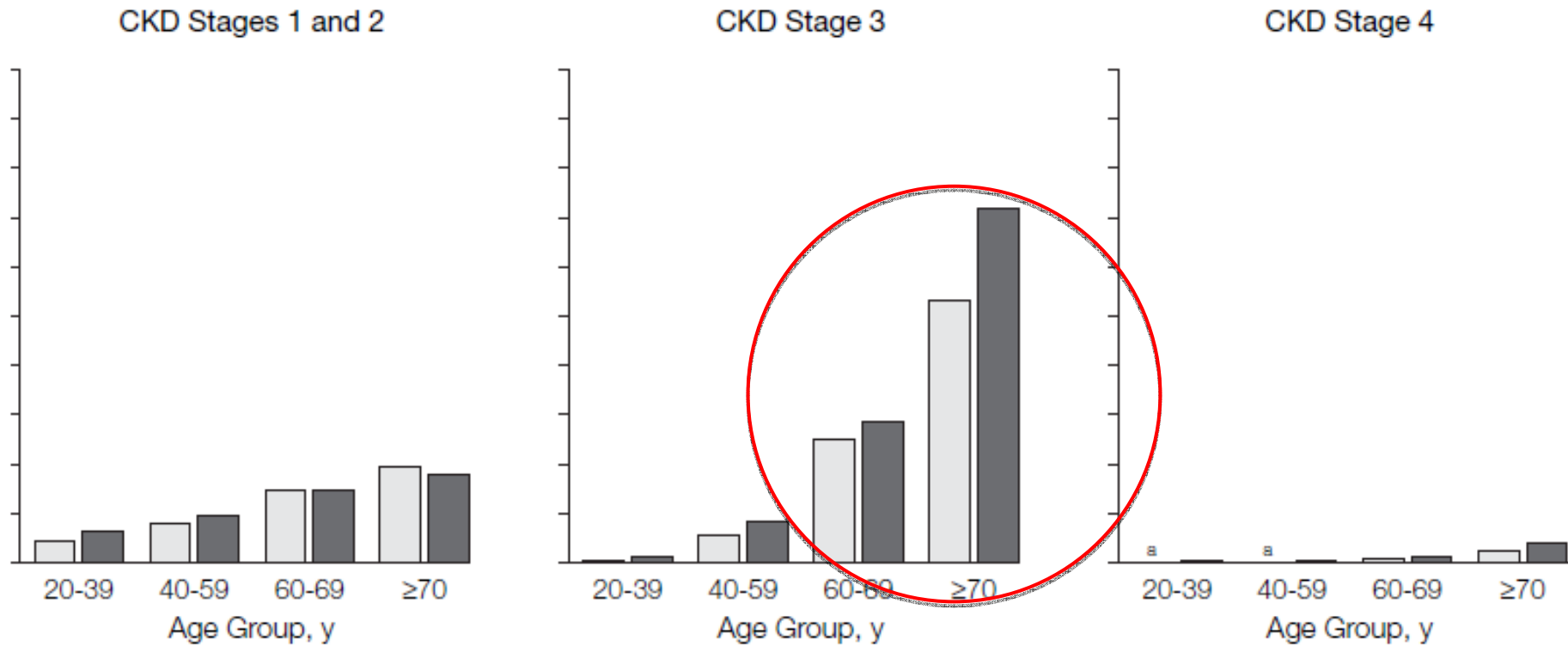
Distribution of NHANES participants with diabetes, congestive heart failure, & markers of CKD, 2005–2010

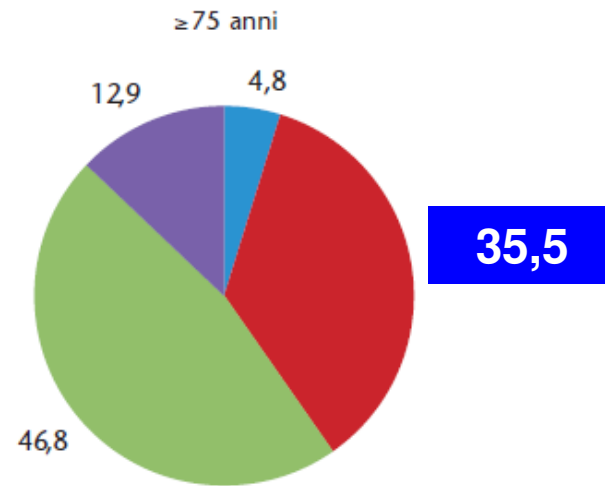
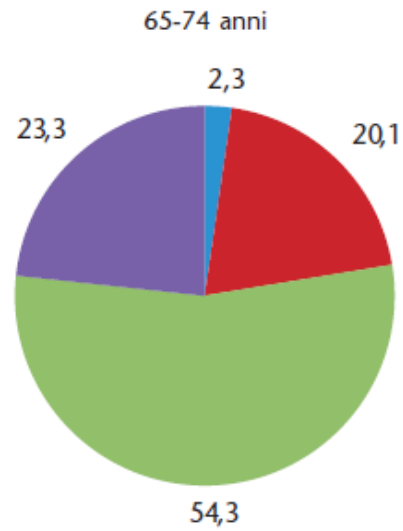
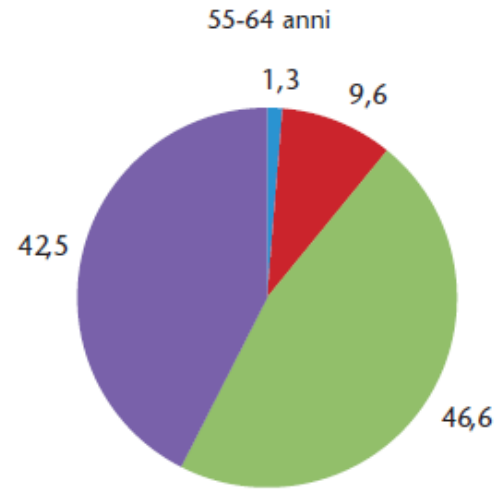
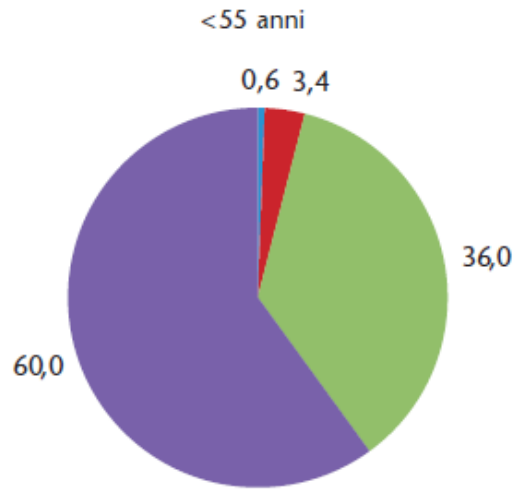
Figure 1.1 (volume 1)



NHANES participants 2005–2010, age 20 & older; eGFR calculated using CKD-EPI equation; urine albumin creatinine ratio (ACR).

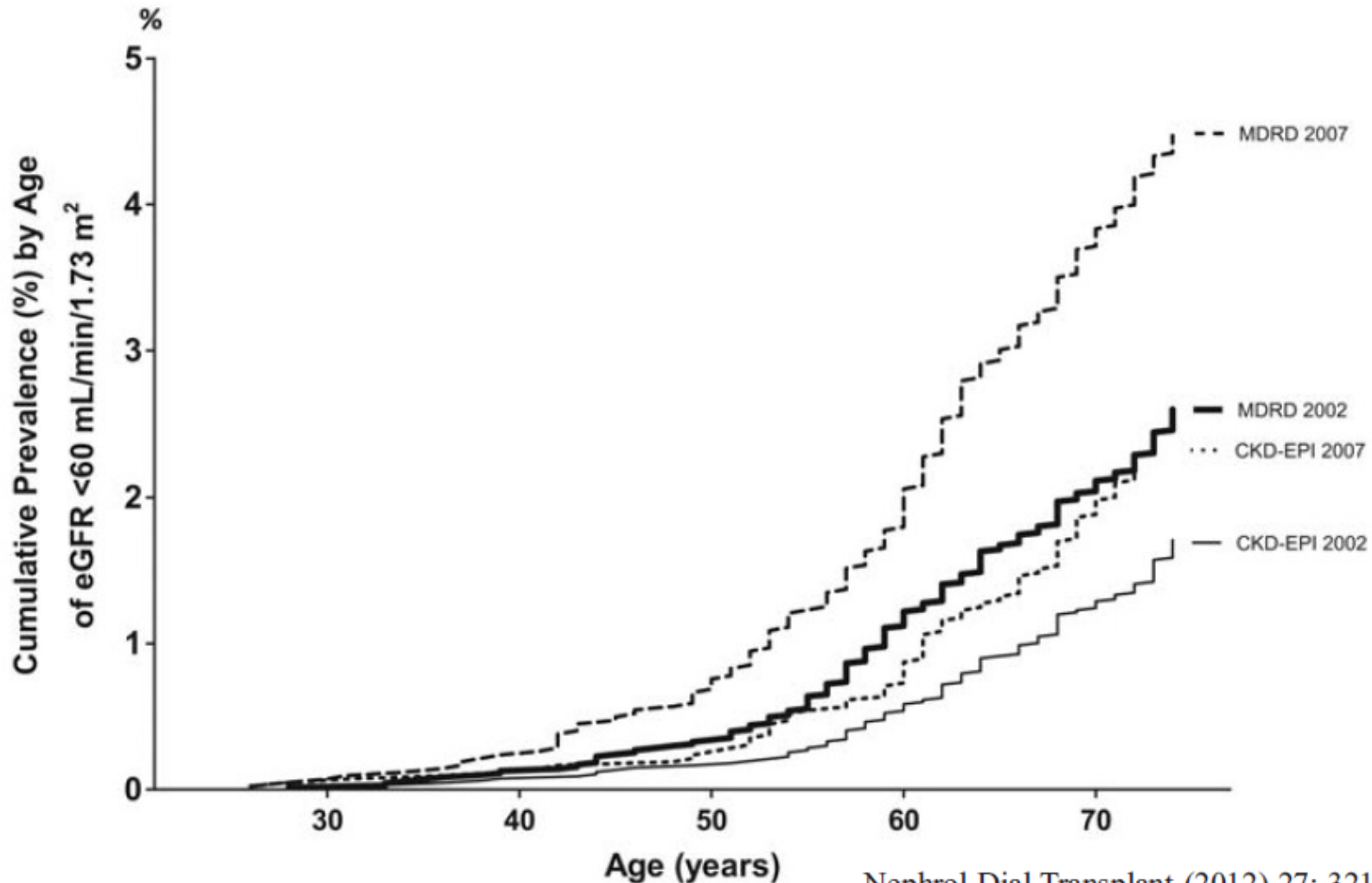
Prevalence of Chronic Kidney Disease in the United States





■ 0-30
 ■ 31-60
 ■ 61-90
 ■ >90 ml/min

Comparison of the MDRD Study and the CKD-EPI Study equations in evaluating trends of estimated kidney function at population level: findings from the National FINRISK Study

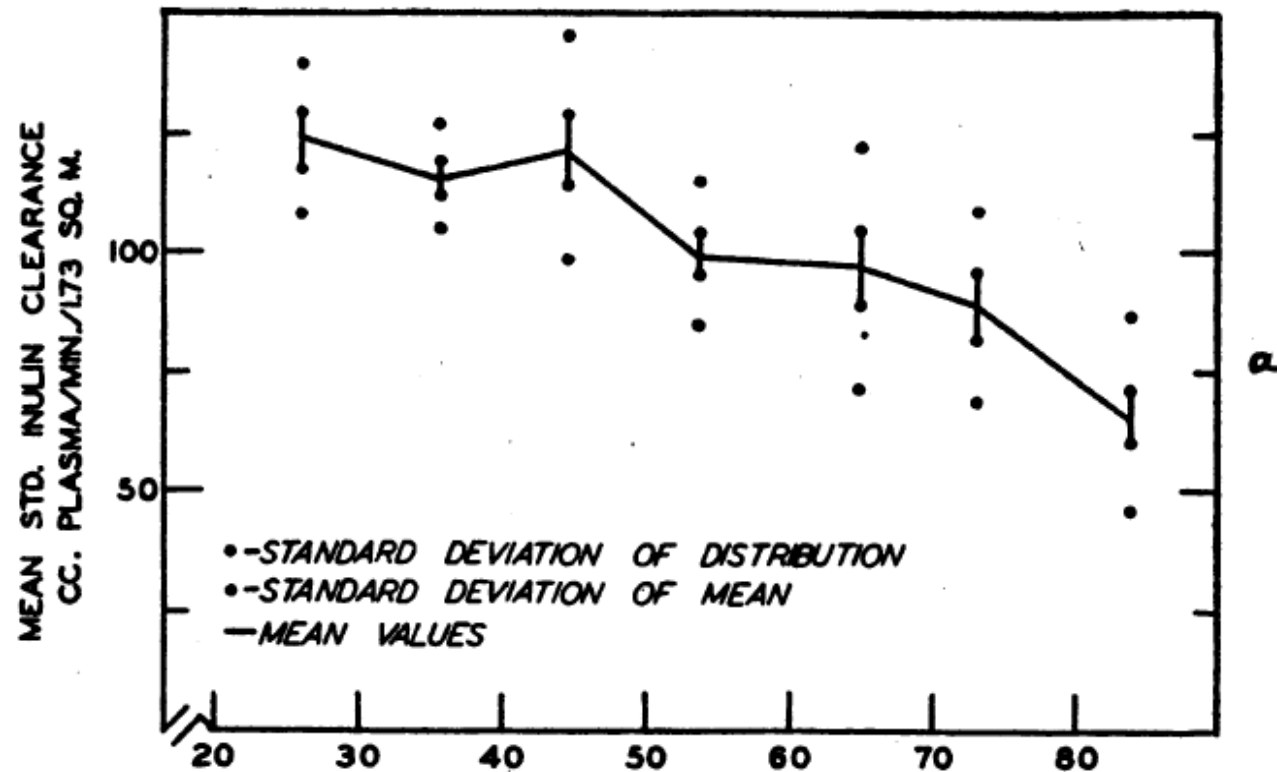


AGE CHANGES IN GLOMERULAR FILTRATION RATE, EFFECTIVE RENAL PLASMA FLOW, AND TUBULAR EXCRETORY CAPACITY IN ADULT MALES

By DEAN F. DAVIES¹ AND NATHAN W. SHOCK

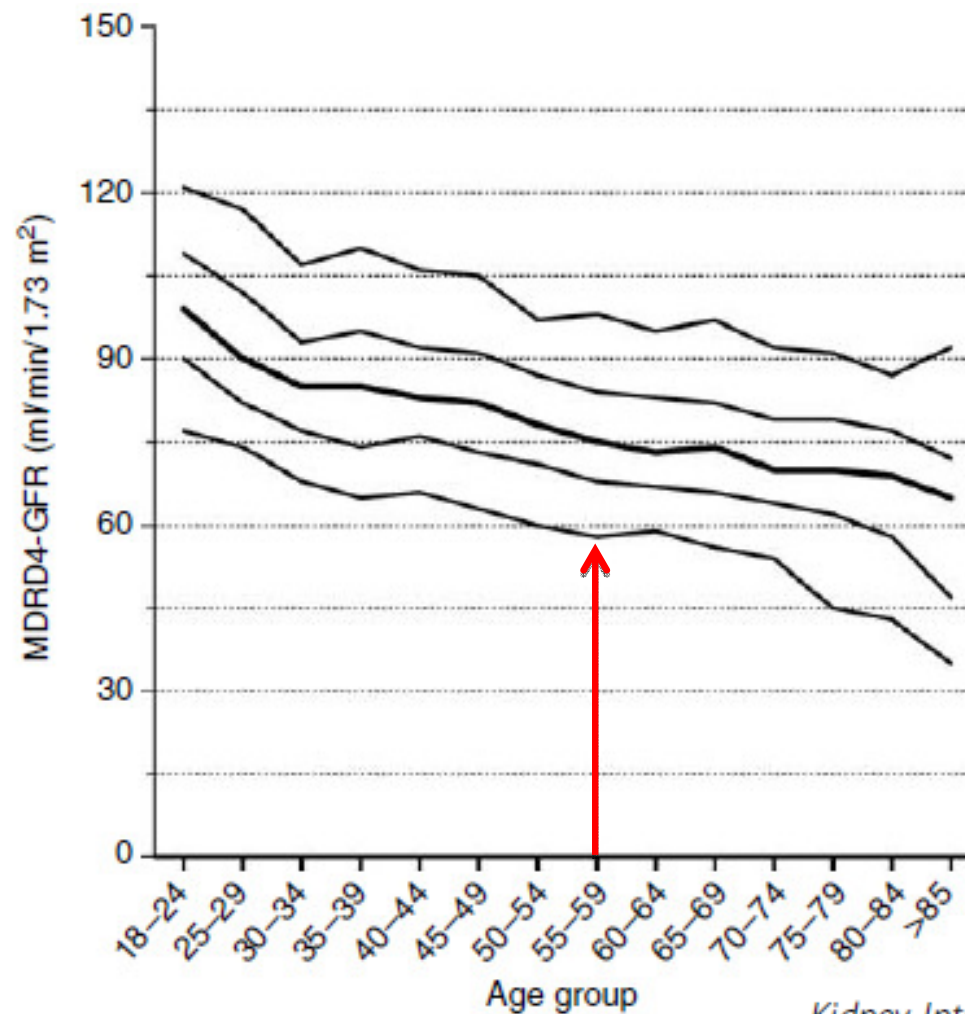
(Submitted for publication October 25, 1949; accepted, December 28, 1949)

AGE CHANGES IN RENAL FUNCTION

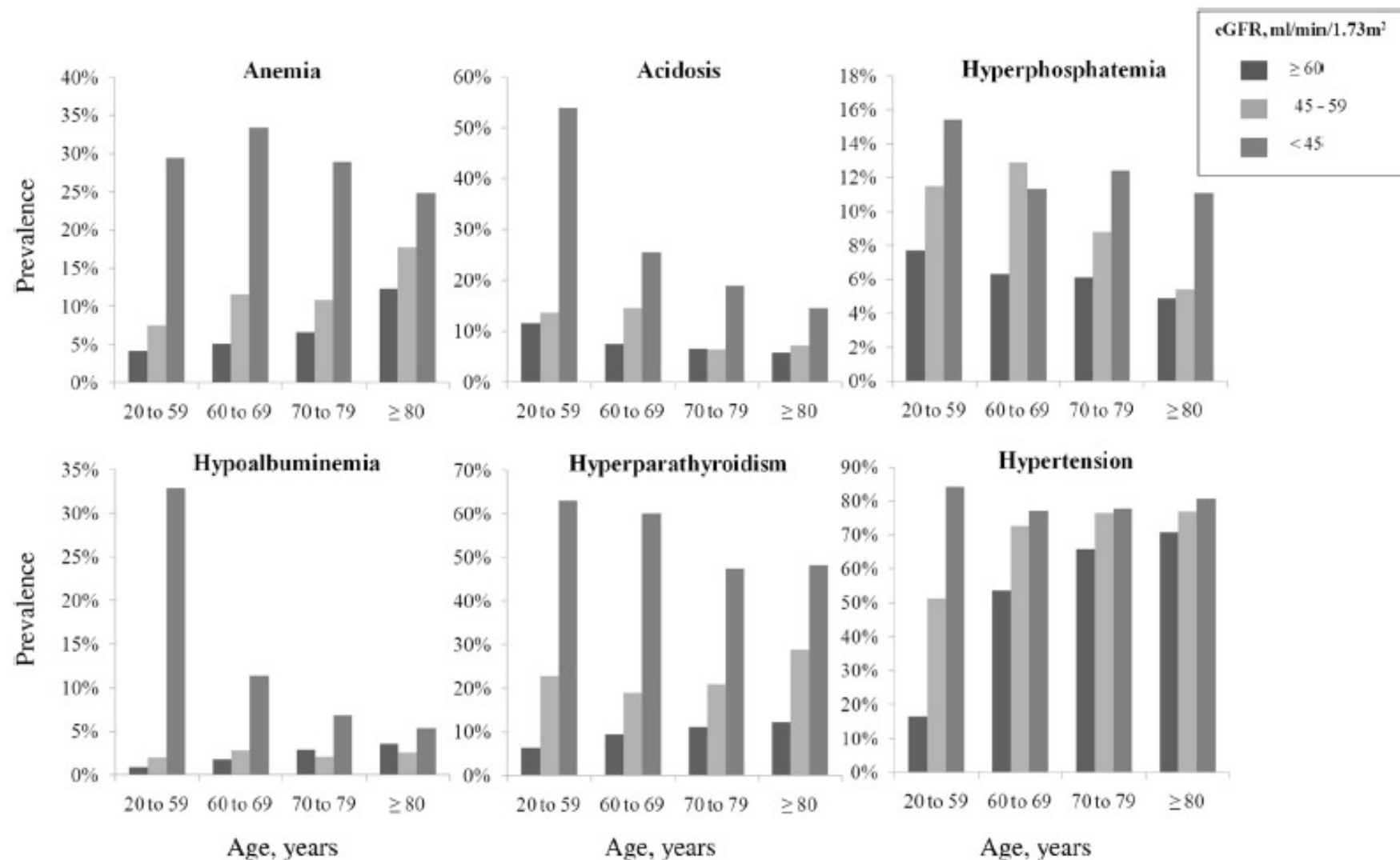


Age- and gender-specific reference values of estimated GFR in Caucasians: The Nijmegen Biomedical Study

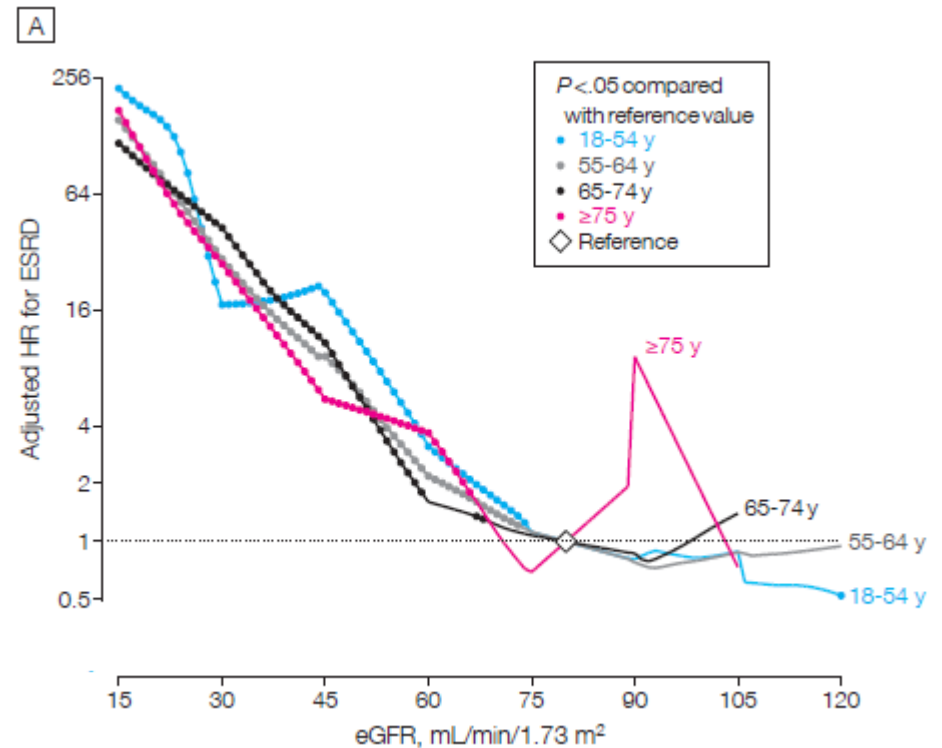
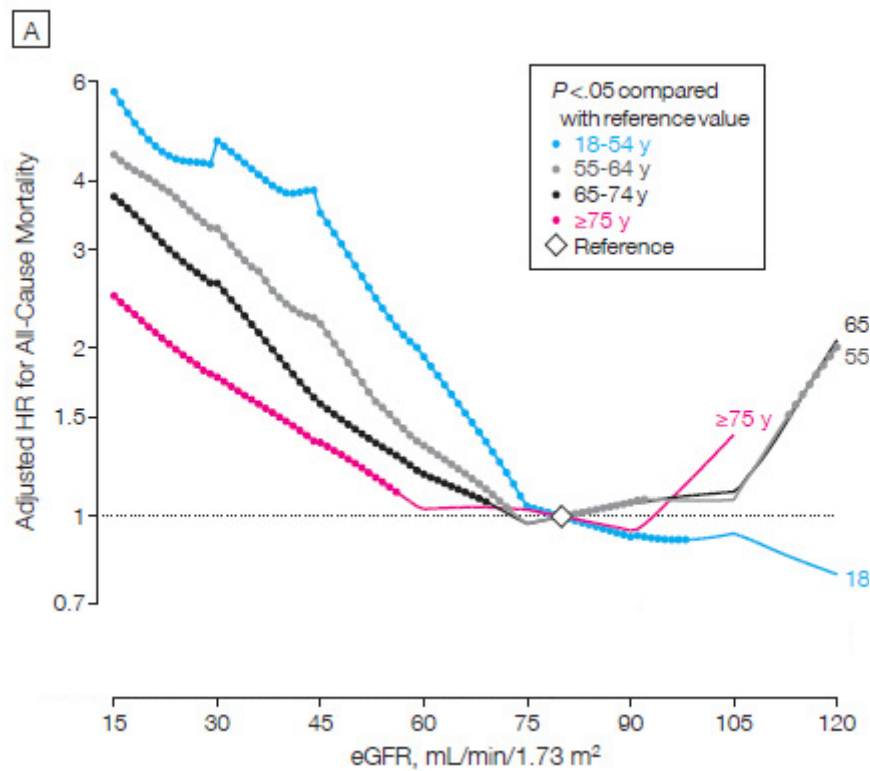
JFM Wetzels¹, LALM Kiemeney², DW Swinkels³, HL Willems³ and M den Heijer^{2,4}



Age-Specific Associations of Reduced Estimated Glomerular Filtration Rate with Concurrent Chronic Kidney Disease Complications



Age and Association of Kidney Measures With Mortality and End-stage Renal Disease



Mortality Risk Stratification in Chronic Kidney Disease: One Size for All Ages?

Age Group (Yr)	eGFR 50 to 59 (<i>n</i> = 38,274)	eGFR 40 to 49 (<i>n</i> = 25,146)
18 to 44 (<i>n</i> = 42,477)	1.24 (0.72 to 2.15)	— ^c
45 to 54 (<i>n</i> = 132,344)	0.86 (0.70 to 1.05)	2.49 (1.21 to 1.84)
55 to 64 (<i>n</i> = 148,669)	0.96 (0.87 to 1.06)	1.28 (1.12 to 1.48)
65 to 74 (<i>n</i> = 167,874)	0.90 (0.85 to 0.96)	1.16 (1.09 to 1.24)
75 to 84 (<i>n</i> = 125,313)	0.94 (0.90 to 0.99)	1.05 (1.00 to 1.10)
85+ (<i>n</i> = 10,377)	0.93 (0.83 to 1.04)	1.04 (0.93 to 1.16)

Chronic Kidney Disease and the Risks of Death, Cardiovascular Events, and Hospitalization

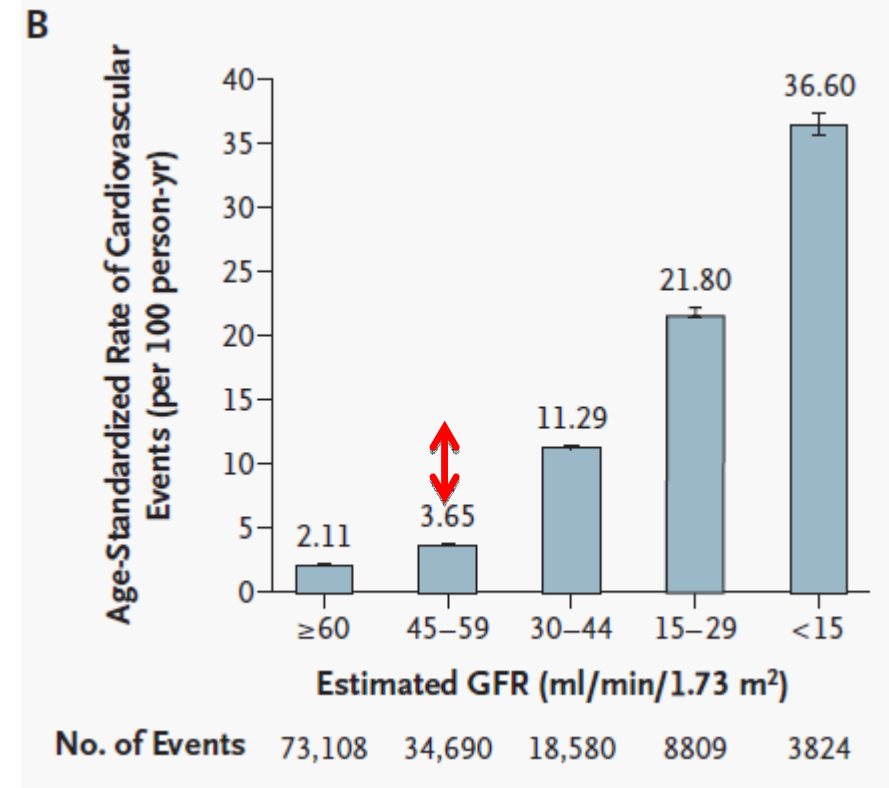
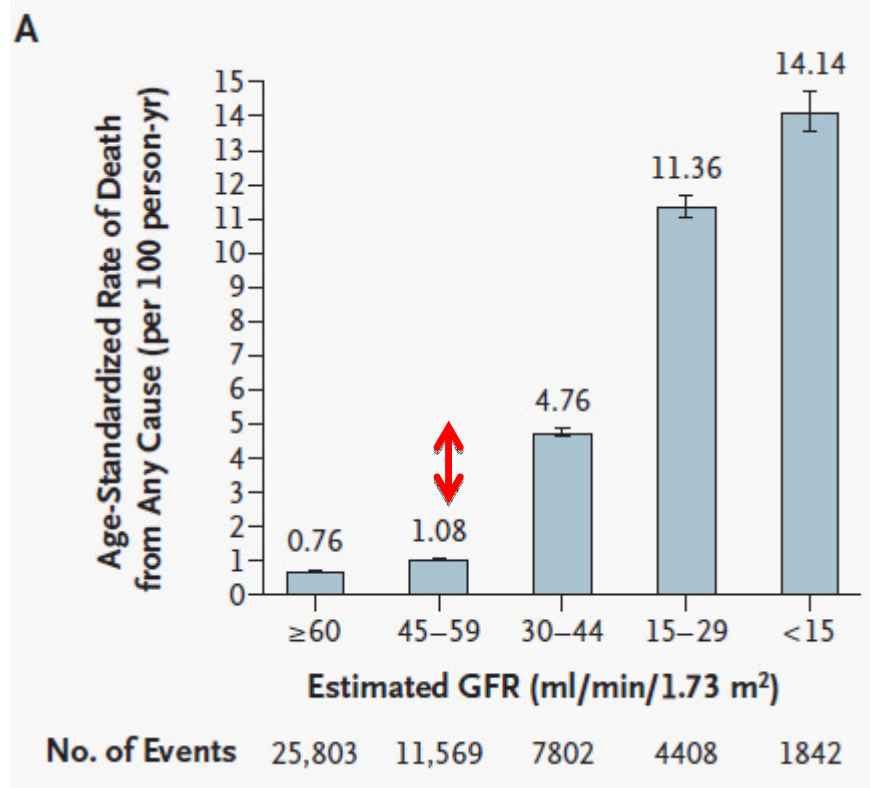
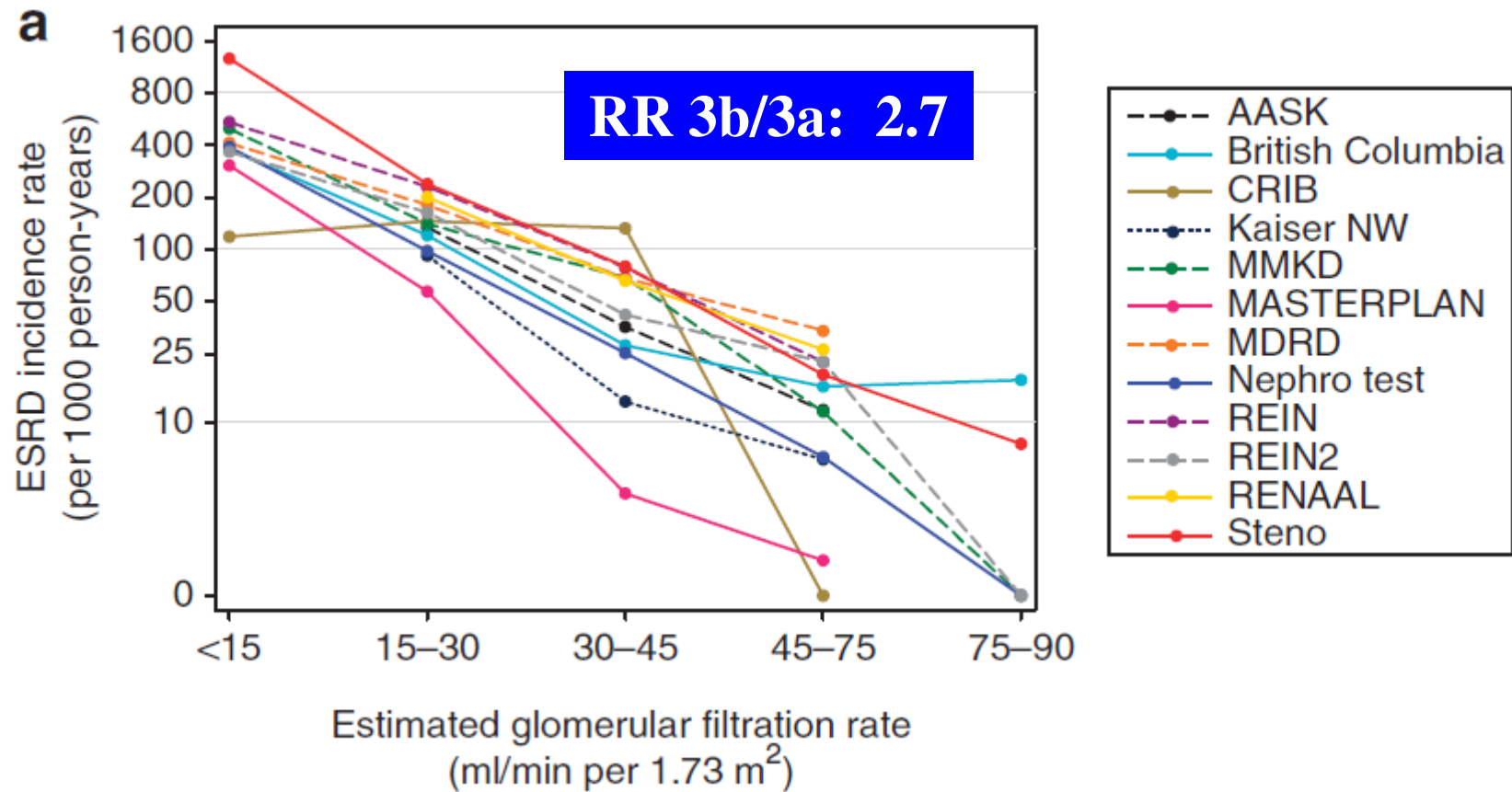




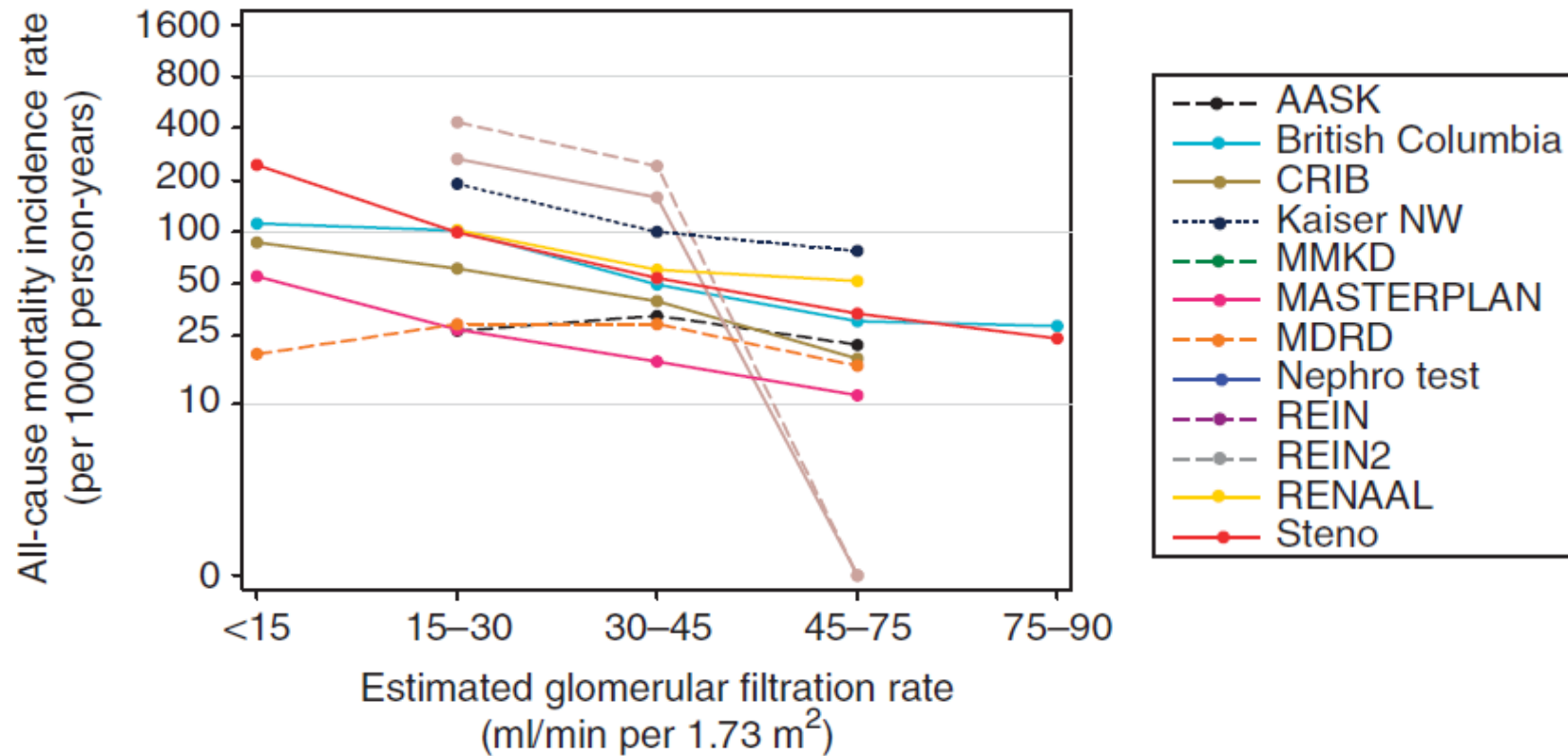
Table 5 | GFR categories in CKD

GFR category	GFR (ml/min/1.73 m ²)	Terms
G1	≥90	Normal or high
G2	60–89	Mildly decreased*
G3a	45–59	Mildly to moderately decreased
G3b	30–44	Moderately to severely decreased
G4	15–29	Severely decreased
G5	<15	Kidney failure

Lower estimated glomerular filtration rate and higher albuminuria are associated with mortality and end-stage renal disease. A collaborative meta-analysis of kidney disease population cohorts



Lower estimated glomerular filtration rate and higher albuminuria are associated with mortality and end-stage renal disease. A collaborative meta-analysis of kidney disease population cohorts



Proteinuria

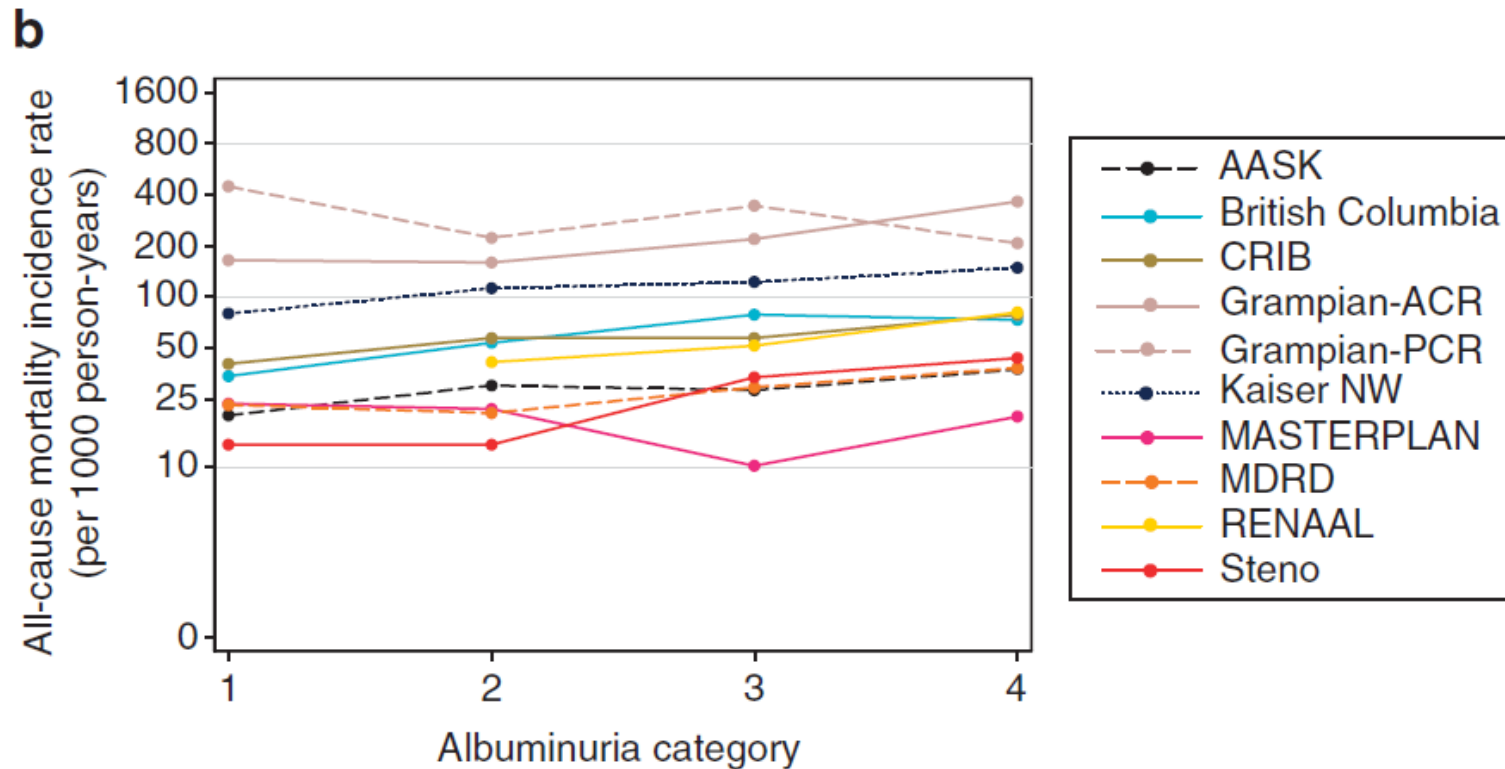
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4	Severe \downarrow in GFR	GFR of 15-29	
5	Kidney failure	GFR < 15 (or dialysis)	D if dialysis

→ Uomo 20 anni, e-GFR 35 mL/min, proteinuria 3 grammi, creatininemia in progressione: **stadio III**.

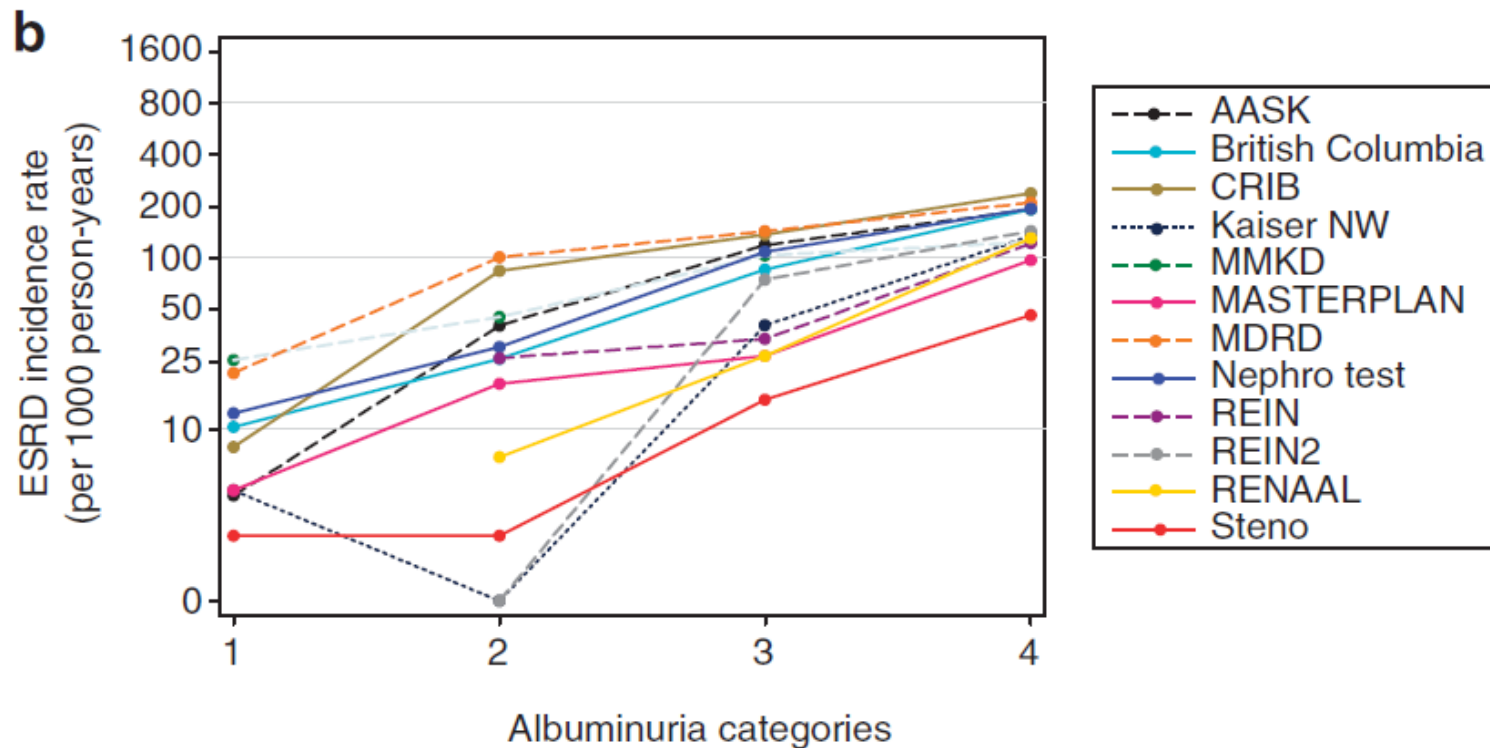
→ Uomo 70 anni, e-GFR 55 mL/min, proteinuria assente, creatininemia stabile: **stadio III**.

Lower estimated glomerular filtration rate and higher albuminuria are associated with mortality and end-stage renal disease. A collaborative meta-analysis of kidney disease population cohorts



Reference	1.46 (1.24, 1.71)	1.80 (1.38, 2.35)	2.26 (1.68, 3.04)
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Lower estimated glomerular filtration rate and higher albuminuria are associated with mortality and end-stage renal disease. A collaborative meta-analysis of kidney disease population cohorts



Reference	2.92 (2.08, 4.10)	7.70 (4.52, 13.10)	15.01 (8.36, 26.95)
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Associations of estimated glomerular filtration rate and albuminuria with mortality and renal failure by sex: a meta-analysis

BMJ 2013;346:f324

End stage renal disease risk by urinary albumin-creatinine ratio (ACR)

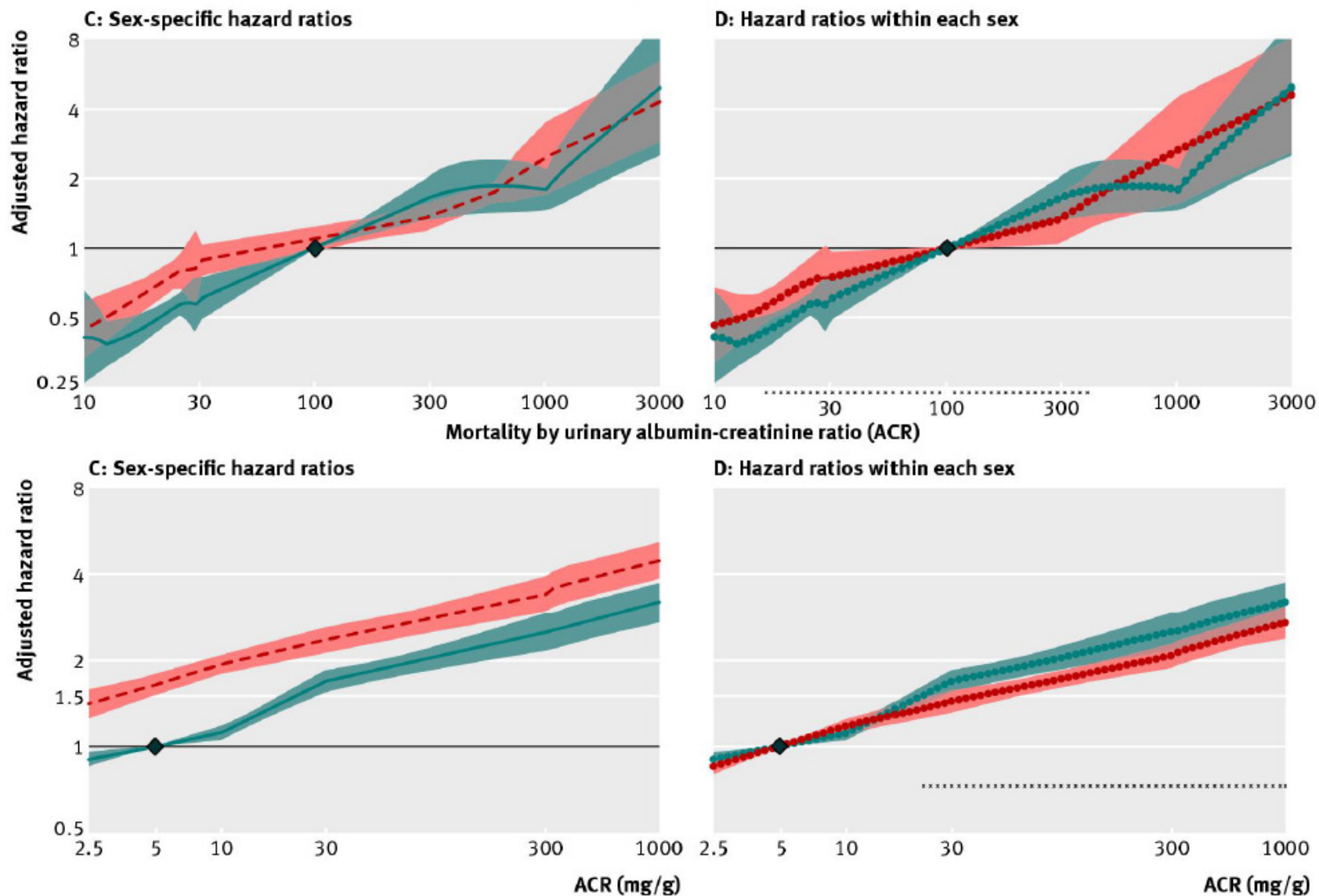
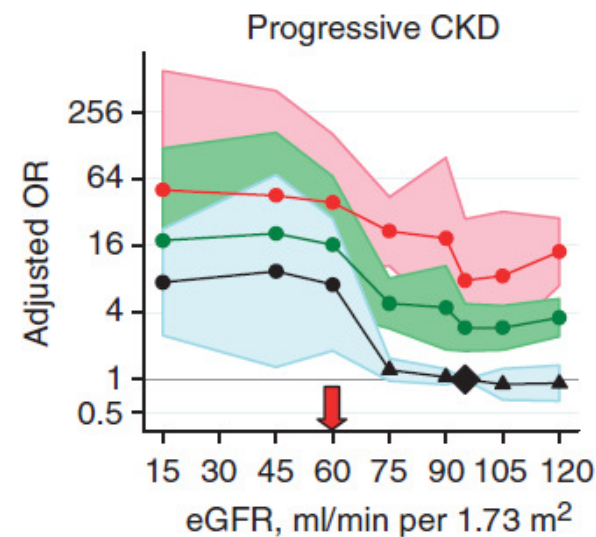
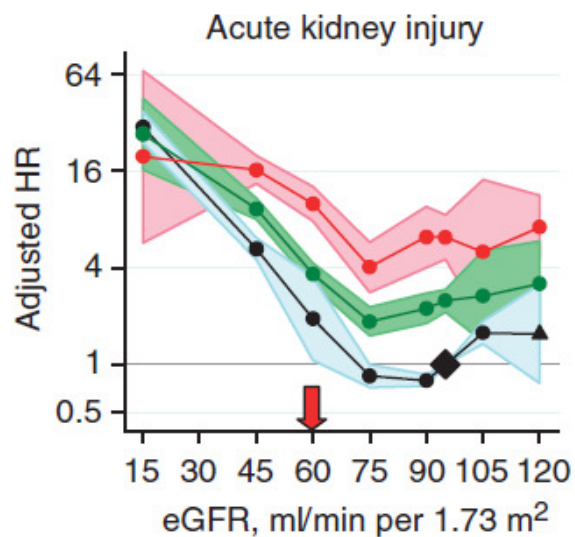
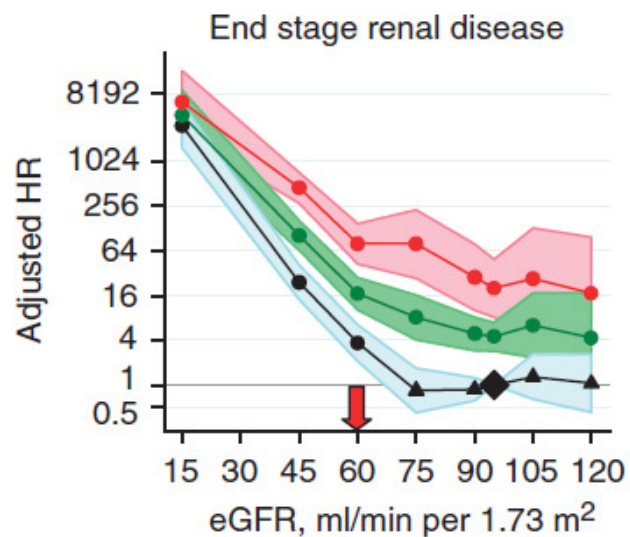
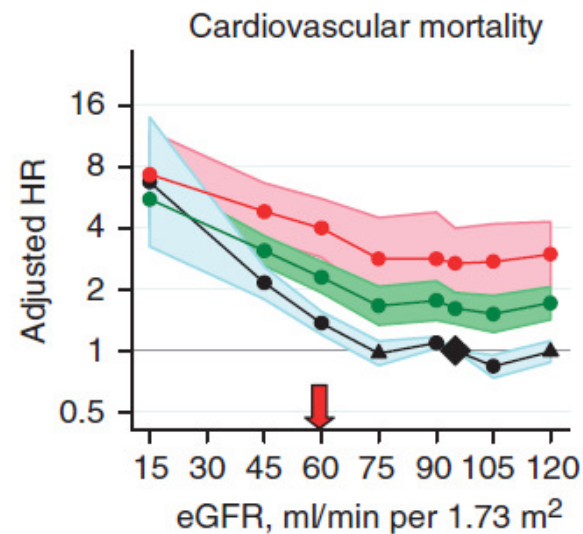
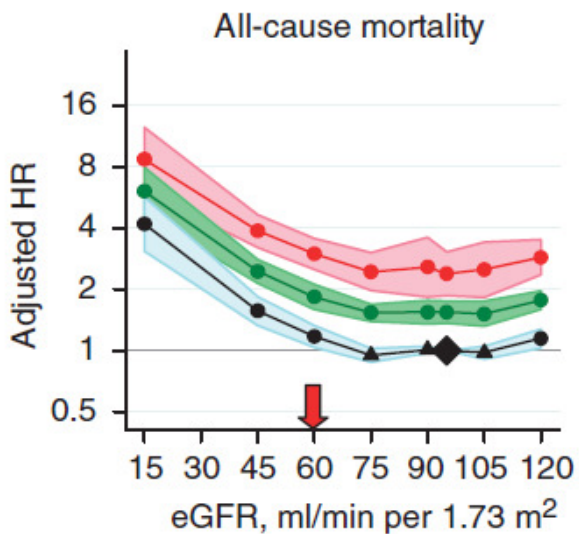




Table 6 | Albuminuria categories in CKD

Category	AER (mg/24 hours)	ACR (approximate equivalent)		Terms
		(mg/mmol)	(mg/g)	
A1	< 30	< 3	< 30	Normal to mildly increased
A2	30-300	3-30	30-300	Moderately increased*
A3	> 300	> 30	> 300	Severely increased**

Summary of relative risks from continuous meta-analysis



- No CKD
- Moderate-risk CKD
- High-risk CKD
- Very high-risk CKD

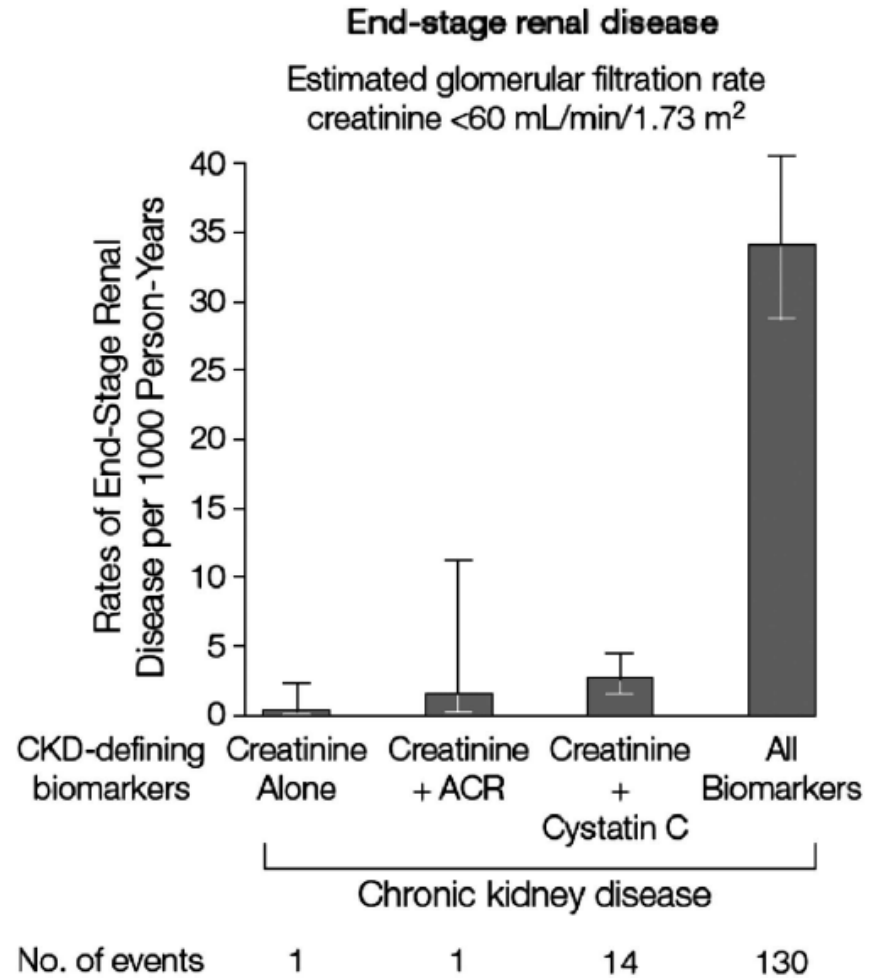
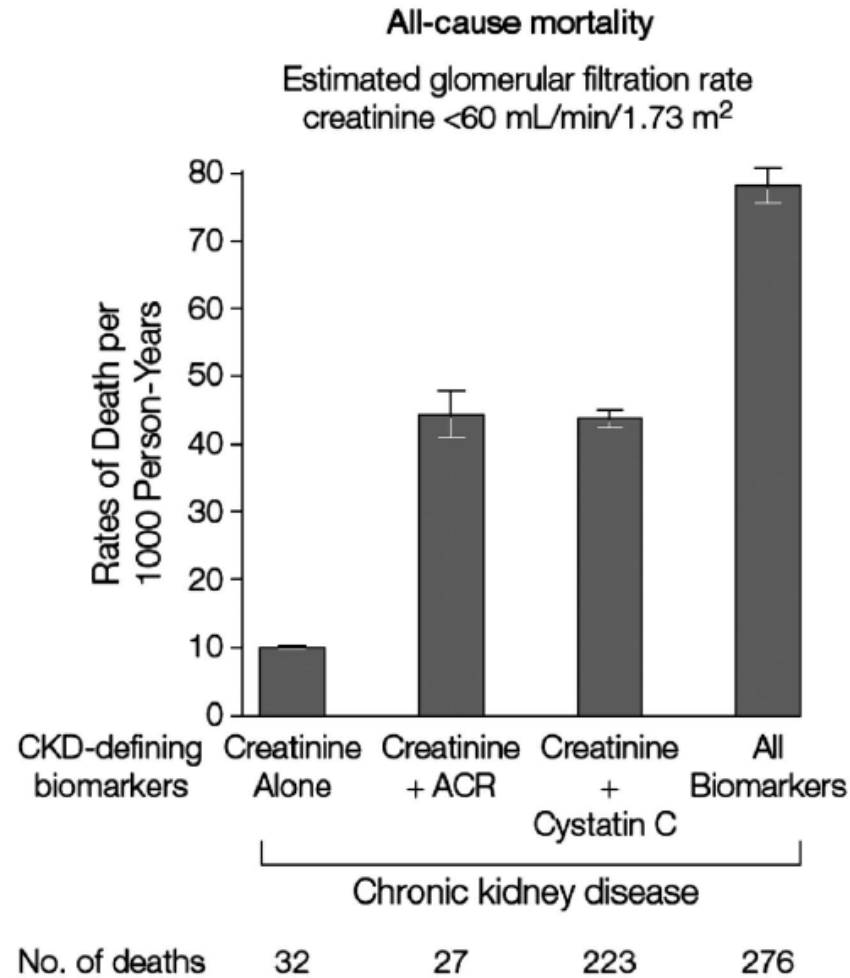
				Albuminuria stages, description, and range (mg/g)				
				A1		A2	A3	
				Optimum and high-normal		High	Very high and nephrotic	
				<10	10-29	30-299	300-1999	≥2000
GFR stages, description, and range (mL/min per 1.73m ²)	G1	High and optimum	>105					
			90-104					
	G2	Mild	75-89					
			60-74					
	G3a	Mild-moderate	45-59					
	G3b	Moderate-severe	30-44					
	G4	Severe	15-29					
G5	Kidney failure	<15						



Cause (?)

Table 4 | Classification* of CKD based on presence or absence of systemic disease and location within the kidney of pathologic-anatomic findings

	Examples of systemic diseases affecting the kidney	Examples of primary kidney diseases (absence of systemic diseases affecting the kidney)
Glomerular diseases	Diabetes, systemic autoimmune diseases, systemic infections, drugs, neoplasia (including amyloidosis)	Diffuse, focal or crescentic proliferative GN; focal and segmental glomerulosclerosis, membranous nephropathy, minimal change disease
Tubulointerstitial diseases	Systemic infections, autoimmune, sarcoidosis, drugs, urate, environmental toxins (lead, aristolochic acid), neoplasia (myeloma)	Urinary-tract infections, stones, obstruction
Vascular diseases	Atherosclerosis, hypertension, ischemia, cholesterol emboli, systemic vasculitis, thrombotic microangiopathy, systemic sclerosis	ANCA-associated renal limited vasculitis, fibromuscular dysplasia
Cystic and congenital diseases	Polycystic kidney disease, Alport syndrome, Fabry disease	Renal dysplasia, medullary cystic disease, podocytopathies



Estimating Glomerular Filtration Rate from Serum Creatinine and Cystatin C

A Bias

