# Estimated GFR in Diabetes 

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## Introduction

- Diabetic nephropathy triad
- Albuminuria rise
- BP rise
- GFR fall
- Abnormal serum creatinine
- Relative late stage in natural history
- Strategies to identify individuals at risk
- Dipstick proteinuria
- Microalbuminuria
- BP


## Introduction

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## MA based strategies - pitfalls

- Uncertain predictive value
- 20-30\% progression (cf. 85-100\% in 1980's)
- Other causes of albuminuria
- Non-albuminuric renal impairment
- Non-diabetic renal disease in diabetes
- $\approx 25 \%$ proven DN \& normoalbuminuria


## GFR

- True GFR measurements unsuitable for mass screening
- Estimated GFR (eGFR)
- From serum creatinine, age, gender, ethnicity...
- Reliable indicators of renal reserve
- Supported by organisations
- National Kidney Foundation
- Renal NSF
- ADA
- (DUK, ABCD)

Table 1—Stages of CKD

| Stage | Description | GFR $\left(\mathrm{ml} / \mathrm{min} / 1.73 \mathrm{~m}^{2}\right.$ <br> body surface area) |
| :--- | :--- | :---: |
| 1 | Kidney damage with normal or increased GFR | $\geq 90$ |
| 2 | Kidney damage with mildly decreased GFR | $60-89$ |
| 3 | Moderately decreased GFR | $30-59$ |
| 4 | Severely decreased GFR | $15-29$ |
| 5 | Kidney failure | $<15$ or dialysis |

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## eGFR equation - C\&G or MDRD

## Aims

- To evaluate renal disease burden in diabetes using eGFR - either by C\&G or MDRD estimate
- To study the clinical utility of eGFR (over and above current markers)


## Methods

- Study design
- cross sectional from district diabetes register
- Study period
- Jan 2002 to June 2003
- MA screening
- spot morning urine ACR ( $3.5 \mathrm{mg} / \mathrm{mmol}$ threshold)
- SPSS 11.5 for statistical analysis


## eGFR equations

- MDRD
$186 \times[\text { Serum } \mathrm{Cr}(\mu \mathrm{mol} / \mathrm{I}) / 88.4]^{-1.154} \times[\text { Age }]^{-0.203} \mathrm{x}$ [0.742 if female] x [1.210 if Black]
- Cockcroft's and Gault's equation (140-age in years) $x$ body weight ( kg ) x K

Serum creatinine ( $\mu \mathrm{mol} / \mathrm{l}$ )
$K=1.23$ for men or 1.04 for women
Correction for BSA of $1.73 \mathrm{~m}^{2}$

## Results

- Total $\mathrm{N}=4548 ; \mathrm{N}$ with eGFR $=4173$

| Age | $60 \pm 14 \mathrm{y}$ |
| :---: | :---: |
| Duration | $12 \pm 9 \mathrm{y}$ |
| BMI | $31 \pm 6 \mathrm{Kg} / \mathrm{m}^{2}$ |
| Males | $57 \%$ |
| Type 2 DM | $78 \%$ |
| Whites/Asians/AfroCarib | $68 \% / 23 \% / 9 \%$ |
| Serum Creatinine | $101 \pm 44 \mu \mathrm{~mol} / \mathrm{I}$ |
| Urine ACR | $1.75 \mathrm{mg} / \mathrm{mmol}$ |

Figure 1a


Figure 1b


## C\&G and MDRD correlation



| Soncordance \& Discordance |  |  |  |  |  |
| :---: | :---: | :---: | :---: | :---: | :---: |
|  | $\begin{aligned} & \text { C\&G } \\ & >90 \end{aligned}$ | $\begin{gathered} \text { C\&G } \\ 90-60 \end{gathered}$ | $\begin{gathered} \text { C\&G } \\ 60-30 \end{gathered}$ | $\begin{aligned} & \text { C\&G } \\ & <30 \end{aligned}$ | Total |
| MDRD $>90$ | $\begin{gathered} 316 \\ (87 \%) \end{gathered}$ | $\begin{gathered} 49 \\ (13 \%) \end{gathered}$ | $\begin{gathered} 0 \\ (0 \%) \end{gathered}$ | $\begin{gathered} 0 \\ (0 \%) \end{gathered}$ | 365 |
| MDRD $90-60$ | $\begin{gathered} 722 \\ (28 \%) \end{gathered}$ | $\begin{gathered} 1557 \\ (61 \%) \end{gathered}$ | $\begin{gathered} 295 \\ (11 \%) \end{gathered}$ | $\begin{gathered} 0 \\ (0 \%) \end{gathered}$ | 2574 |
| $\begin{gathered} \text { MDRD } \\ 60-30 \end{gathered}$ | $\begin{gathered} 10 \\ (1 \%) \end{gathered}$ | $\begin{gathered} 315 \\ (28 \%) \end{gathered}$ | $\begin{gathered} 795 \\ (70 \%) \end{gathered}$ | $\begin{gathered} 22 \\ (2 \%) \end{gathered}$ | 1142 |
| MDRD <30 | $\begin{gathered} 0 \\ (0 \%) \end{gathered}$ | $\begin{gathered} 0 \\ (0 \%) \end{gathered}$ | $\begin{gathered} 33 \\ (36 \%) \end{gathered}$ | $\begin{gathered} 59 \\ (64 \%) \end{gathered}$ | 92 |

## Concordance \& Discordance <br> 

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## 65\% Green

## Concordance \& Discordance

|  | C\&G <br> $>90$ | C\&G <br> $90-60$ | C\&G <br> $60-30$ | C\&G <br> $<30$ | Total |
| :---: | :---: | :---: | :---: | :---: | :---: |
| MDRD <br> $>90$ | 316 <br> $(87 \%)$ | 49 | 0 | 0 | 365 |
| MDRD | $72 \%$ | $(0 \%)$ | $(0 \%)$ |  |  |
| $90-60$ | $(28 \%)$ | 1557 | $\mathbf{2 9 5}$ | 0 | 2574 |
| MDRD | $\mathbf{1 0}$ | $\mathbf{3 1 5}$ | $\mathbf{( 1 1 \% )}$ | $(0 \%)$ |  |
| $60-30$ | $\mathbf{( 1 \% )}$ | $\mathbf{( 2 8 \% )}$ | $(70 \%)$ | 22 | 1142 |
| MDRD | 0 | 0 | 33 | 59 | 92 |
| 30 | $(0 \%)$ | $(0 \%)$ | $(36 \%)$ | $(64 \%)$ |  |

## 65\% Green;

## Concordance \& Discordance

|  | C\&G <br> $>90$ | C\&G <br> $90-60$ | C\&G <br> $60-30$ | C\&G <br> $<30$ | Total |
| :---: | :---: | :---: | :---: | :---: | :---: |
| MDRD <br> $>90$ | 316 <br> $(87 \%)$ | 49 | 0 | 0 | 365 |
| MDRD | $72 \%$ | 1357 | $(0 \%)$ | $(0 \%)$ |  |
| $90-60$ | $(28 \%)$ | $(61 \%)$ | $(11 \%)$ | 0 | 2574 |
| MDRD | 10 | 315 | 795 | 22 | 1142 |
| $60-30$ | $(1 \%)$ | $(28 \%)$ | $(70 \%)$ | $(2 \%)$ |  |
| MDRD <br> $<30$ | 0 | 0 | 33 | 59 | 92 |

## 65\% Green; <br> \& 15\% Red



## Renal risk markers in those with serious discordance

|  | C\&G<60 <br> MDRD>60 <br> N=295 | MDRD<60 <br> C\&G>60 <br> $\mathbf{N = 3 2 5}$ |
| :---: | :---: | :---: |
| Abnormal serum <br> Creatinine | $28(10 \%)$ | $51(16 \%)$ |
| Abnormal urine <br> ACR | $90(31 \%)$ | $112(35 \%)$ |
| Abnormal <br> creatinine or ACR | $107(36 \%)$ | $136(42 \%)$ |

## Study summary

- Renal disease burden was different depending on the eGFR equation used
- Full concordance observed in 65\%
- Serious discordance in 15\%
- The majority with serious discordance had normal levels of other renal markers
- Relying entirely on eGFR to flag their risk
- What does low eGFR really mean?


## eGFR, RRT \& Mortality

| N=28,000 | Stage 2 | Stage 3 | Stage 4 |
| :--- | :--- | :--- | :--- |
| RRT <br> (within 3yr) | $1.1 \%$ | $1.3 \%$ | $19.9 \%$ |
| Mortality | $19.5 \%$ | $24.3 \%$ | $45.7 \%$ |

Keith et al, Arch Intern Med 2004

## eGFR, mortality \& CVS events



$\begin{array}{llllll}\text { No. of Events } & 73,108 & 34,690 & 18,580 & 8809 & 3824\end{array}$

## Discussion - role of eGFR

- Renal progression indicator
- Predictor of mortality \& CVS events
- Role in predicting safety of Metformin?
- Early and inexpensive identification of risk individuals
- No data to support intervention solely based on eGFR
- Lack of standardization of creatinine across labs
- Validation in diabetes lacking
- Exaggerates risk in the very old?


## Conclusion

- eGFR may have an additional role in renal and vascular risk prediction
- Need for a single equation of choice
- Clarity
- Uniformity of practice

