Electronic Tools to for Creatinine Clearance Calculation to Support Dosing of Direct-Acting Oral Anticoagulants (DOACs)

Summary:

- This is an updated to the previous version of this BSW document to cover the NEW renal disease calculator included within SystmOne in October 2023. The old SystmOne renal disease calculator will be removed from SystmOne clinical tools in subsequent SystmOne maintenance updates.
- Always use creatinine clearance for direct-acting oral anticoagulants (DOAC) dose calculations and <u>NOT</u> eGFR as recommended by the MHRA and SPCs for relevant DOACs. Use of eGFR for dosing of DOACs is known to increase risk of bleeding events as a consequence of overestimating renal function.
- Use either the SystmOne renal disease calculator or MD+Calc for calculating creatinine clearance as per your current practice.
- Based on several papers (Winter et al 2012 and Brown et al 2013) and local expert opinions, BSW ICB support an adjustment to the Cockcroft-Gault equation based on patient's BMI, as it appears to become less accurate in weight extremes (underweight and particularly overweight/obesity). Based on local experts' consensus, adjustments and estimates should be considered as follow:

_	Body Mass Index (kg/m ²)	Adjustments and Estimates
Underweight or Normal weight or		
Overweight	BMI < 30	Calculate CrCl by using actual body weight
Obese	BMI ≥30	Calculate CrCl by using adjusted body weight

- When interpreting creatine clearance, ensure the renal function is in a steady state and an up-to-date serum creatinine are documented. Take into account the trend of creatine clearance changes rather than point estimated value while making any relevant dosage adjustment.
- When interpreting BMI, ensure an up-to-date weight (in last 12 months) and height (in last 5 years) are documented. Take into account any other unintentional rapid weight changes, sign of acute stress or a serious illness that may require most accurate available measurements.
- Caution in clinical judgement and extra interpretation may be require when the range falls within a dose adjustment boundary. Specialist Anticoagulant advice for these patients should be obtained from our local hospitals:
 - **GWH**: gwh.anticoag.clinic@nhs.net or sarah.bond6@nhs.net Tel: 01793 60434
 - **RUH**: ruh-tr.AnticoagulationTeam@nhs.net or nathan.hutchinson-jones@nhs.net or via Cinapsis
 - SFT: nicolamcquaid@nhs.net or sft.anticoagulation.service@nhs.net

Background

Estimated glomerular filtration rate (eGFR) and creatinine clearance (CrCl) are two estimates of renal function available to prescribers. Clinical laboratories routinely report renal function in adults based on eGFR normalised to a body surface area of 1.73 m². For most drugs and most situations, eGFR is an acceptable estimate of renal function.

However, eGFR can overestimate renal function compared with CrCl in some patient groups or clinical situations. This overestimation can result in patients receiving higher than recommended doses of their medicine in relation to their renal function. CrCl should be considered for dosage adjustment of medicines that are substantially renally excreted and have a narrow therapeutic index. In particular, CrCl should always be used to guide dose adjustment for DOACs.

Practical consideration when utilising the NEW SystmOne Renal Disease Calculator and MD+Calc

There are currently NO national consensus on how to adjust the Cockcroft-Gault equation based on a patient's weight. The BNF recommends using ideal body weight to calculate the CrCl. Except in underweight patients where actual body weight should be used. The MHRA guideline recommends that we should be calculating CrCl to assess renal function and potential dose adjustments in patients taking DOACs and not using the eGFR. It indicates MD+Calc as a suitable tool for calculating CrCl but it does so in a way not to exclude the use of other tools. It makes no suggestion to what weight should be used in the calculation other than to say that 'MD+Calc offers the ability to use adjusted body weight, ideal body weight or actual body weight as appropriate'. See <u>MHRA guidance</u> for more details.

Based on several papers (Winter et al 2012 and Brown et al 2013) and local expert opinions, BSW ICB support an adjustment to the Cockcroft-Gault equation based on a patient's BMI, as it appears to become less accurate in weight extremes (underweight and particularly overweight/obesity). Based on local experts' consensus, adjustments and estimates should be considered as follow:

Underweight, BMI < 18.5 kg/m²

🏋 Renal Disease Calculations 🛛 🗙				
Parameters				
Age	78 🗦 years			
Sex	O Male Female			
Height	1.7 📩 m Latest value (1.7 m) recorded on 01 Nov 2023			
Weight	45 📩 kg Latest value (45.0 kg) recorded on 01 Nov 2023	3		
Serum Creatinine	120 🕂 umoVL			
Results				
BMI 15.57 Kg/m ² is classed as Underweight				
Cockcroft-Gault Formula:				
Creatinine clearan	ce using ideal weight 33.3 ml/min			
Creatinine clearan	ce using actual weight 24.4 ml/min			
Creatinine clearance using adjusted weight 29.7 ml/min				
	About Reset Close			

Based on local experts' consensus, for underweight patient (BMI <18.5 kg/m²) consider calculating CrCl by using **actual body weight.**

	Female	Male		
Age	78	years		
Weight	45	kg 与		
Creatinine	120	µmol/L 띀		
The Cockcroft-Gault Equation may be inaccurate depending on a patient's body weight and BMI; by providing additional height, we can calculate <u>BMI</u> and provide a modified estimate and range.				
Height	170	cm 띀		

MD+Calc presents the result of CrCl for underweight patient, using **actual body weight**.

Normal weight or overweight 30 kg/m² > BMI > 18.5 kg/m²

🍸 Renal Disease Calculations 🛛 🗡		
Parameters		
Age 78	years	
Sex 🔿	Male 💿 Female	
Height 1.7	m Latest value (1.7 m) recorded on 21 Nov 2023	
Weight 60	kg Latest value (60.0 kg) recorded on 21 Nov 2023	
Serum Creatinine 120) 🕆 umol/L	
Results BMI 20.76 Kg/m² is classed as Normal Cockcroft-Gault Formula:		
Creatinine clearance us	sing ideal weight 33.3 ml/min	
Creatinine clearance us	sing <mark>actual weig</mark> ht 32.5 ml/min	
Creatinine clearance using adjusted weight 33 ml/min		
	About Reset Close	

Based on local experts' consensus, for normal weight or overweight patient (BMI 18.5- 29.9 kg/m²) consider calculating CrCl by using **actual body weight**

Overweight/Obese, BMI ≥30 kg/m²

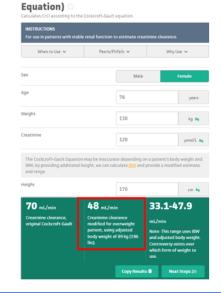
🝸 Renal Disease Calculations 🛛 🗙				
Parameters				
Age	78 🔆 years			
Sex	🔿 Male 🖲 Female			
Height	1.7 m Latest value (1.7 m) recorded on 01 Nov 2023			
Weight	130 🗦 kg Latest value (130.0 kg) recorded on 01 Nov 2023			
Serum Creatinine	120 📫 umol/L			
Results				
BMI 44.98 Kg/m² is classed as Obese II				
Cockcroft-Gault Formula:				
Creatinine clearan	ce using ideal weight 33.3 ml/min			
Creatinine clearance using actual weight 70.5 ml/min				
Creatinine clearance using adjusted weight 48.2 ml/min				
	About Reset Close			

Based on local experts' consensus, for overweight/obese patients (BMI ≥30 kg/m²) consider calculating CrCl by using **adjusted body weight.**

Creatinine Clearance (Cockcroft-Gault Equation) ☆

INSTRUCTIONS For use in patients with stable renal function to estimate creationic dearance. When to Use v Pearlu/Pitfails v Why Use v Sex Male remain Age 78 years Male remain Age 78 years Weight 60 kg 4 Creationice 120 ymovil. 4g The Cockcroft-Gault Equation may be inaccurate depending on a pasient's biody weight and Bible, by providing additional height, we can calculate Male diprovide a modified estimate and range. Weight 100 cm 4g 32 mL/min Creatione clearance, original Cockcroft-Gault Stag (133) Bibl. Copy Besults 8 March 2000 provide a modified estimate Copy Besults 8 March 2000 provide a modified estimate March 2000 provide patients of the stage dearance for sing deal Dody weight of Stag (133) Bibl. Copy Besults 8 March 2000 provide patients of the stage dearance for which form of weight to use. Copy Besults 8 March 2000 provide patients of the stage dearance for sing deal Dody weight of Stage (133) Bibl. Copy Besults 8 March 2000 provide patients of the stage dearance for sing deal Dody weight of Stage (133) Bibl. Copy Besults 8 March 2000 provide patients of the stage dearance for sing deal Dody weight of Stage (133) Bibl. Copy Besults 8 March 2000 provide patients of the stage dearance for sing deal Dody weight of Stage (133) Bibl. Copy Besults 8 March 2000 provide patients of the stage dearance for sing deal Dody weight of Stage (133) Bibl. Copy Besults 8 March 2000 provide patients of the stage dearance for sing deal Dody weight of Stage dearance for stage dearance for sings deal Dody weight of Stage dearance for stage dearance for sings deal Dody weight of Stage dearance for stage dearance for sings deal Dody weight of Stage dearance for stage de

MD+Calc presents the result of CrCl for a normal weight patient, using **actual** and ideal body weight as a 'value range' due to lack of consensus over which form of weight to use.



Creatinine Clearance (Cockcroft-Gault

MD+Calc presents the result of CrCl for obese patient, calculated using **adjusted body weight**, and also presents a value range due to lack of consensus over which form of weight to use.

Based on local experts' consensus, for MD+Calc presents the result of CrC

Exercise caution in clinical judgement and interpretation when the range falls within a dosage adjustment boundary. For instance, in the obese scenario, we observed that the CrCl; results by actual weight and adjusted weight falls into a different Edoxaban BNF dosage boundary for prophylaxis of stroke. Based on local experts' consensus, use the result generated by use of the adjusted weight or seek advice from your local Anticoagulant Specialist in the event of any uncertainty regarding which dose is best to use.

Specialist Anticoagulant advice for these patients can also be obtained from our local hospitals:

- **GWH**: gwh.anticoag.clinic@nhs.net or sarah.bond6@nhs.net Tel: 01793 60434
- **RUH**: ruh-tr.AnticoagulationTeam@nhs.net or nathan.hutchinson-jones@nhs.net or via Cinapsis
- SFT: nicolamcquaid@nhs.net or sft.anticoagulation.service@nhs.net

Reference Sources

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Brown, D.L., Masselink, A.J. and Lalla, C.D. (2013). Functional Range of Creatinine Clearance for Renal Drug Dosing: A Practical Solution to the Controversy of Which Weight to Use in the Cockcroft-Gault Equation. *Annals of Pharmacotherapy*, 47(7-8), pp.1039–1044. doi:https://doi.org/10.1345/aph.1s176.

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