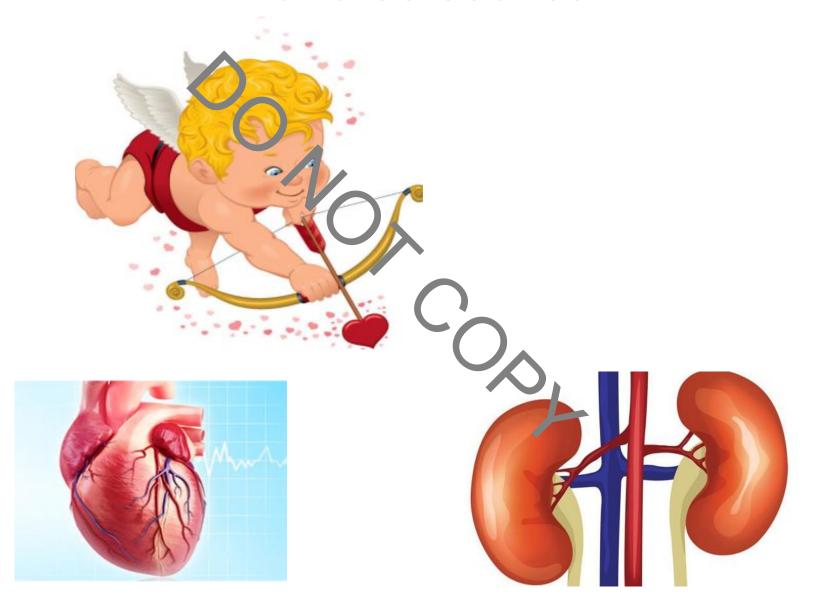
What is more important: the Kidneys or the Heart?

Philip A Kalra
Professor of Nephrology,
Salford Royal Hospital and University of
Manchester, UK

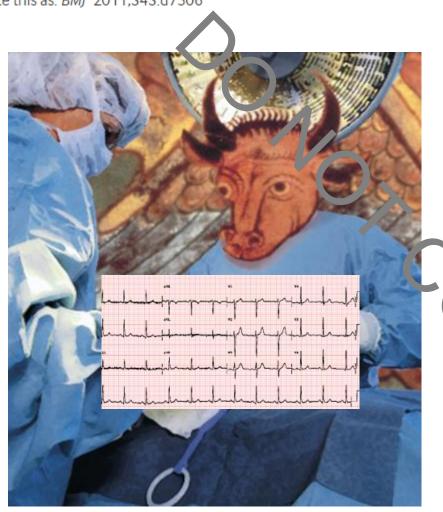
Initial disclosures



Orthopaedic surgeons: as strong as an ox and almost twice as clever? Multicentre prospective comparative study

BMJ 2011; 343 doi: https://doi.org/10.1136/bmj.d7506 (Published 15 December 2011)

Cite this as: BMJ 2011;343:d7506



...and defining the double blind clinical trial

Is there really any doubt about which organ is more important!

How amazing are the Kidneys..!

by Dr. Vall | Apr 21, 2020, | How amazing are the Kidneys...!

Kidneys Are Amazing Organs

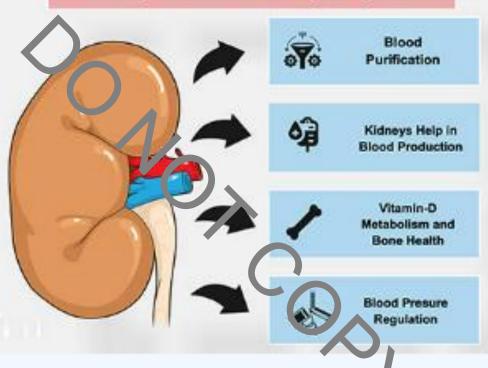
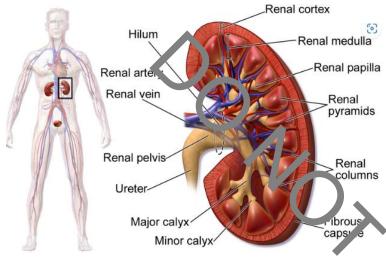


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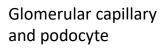
- Each Kidney = One million tiny Nephrons
- Blood is filtered 400 times in a Day
- Kidneys are the True Multitaskers
- Kidneys regulate the Blood Pressure Regulation
- Four Common Culprits for Kidney Failure
- · Protect the amazing Organs

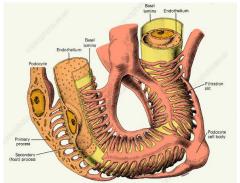


Normal length 8 – 13 cm

Weight ~ 160 gm

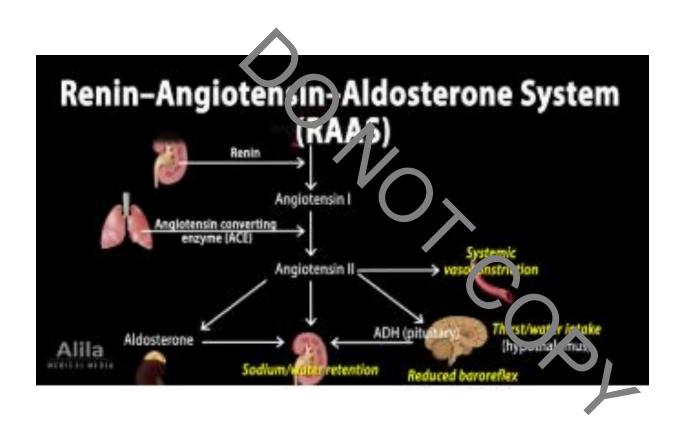
Blood flow 1 – 1.25 litres/min (25% of cardiac output)







Functions of the kidneys



Big role in autoregulation:

- blood pressure control
- salt and fluid balance

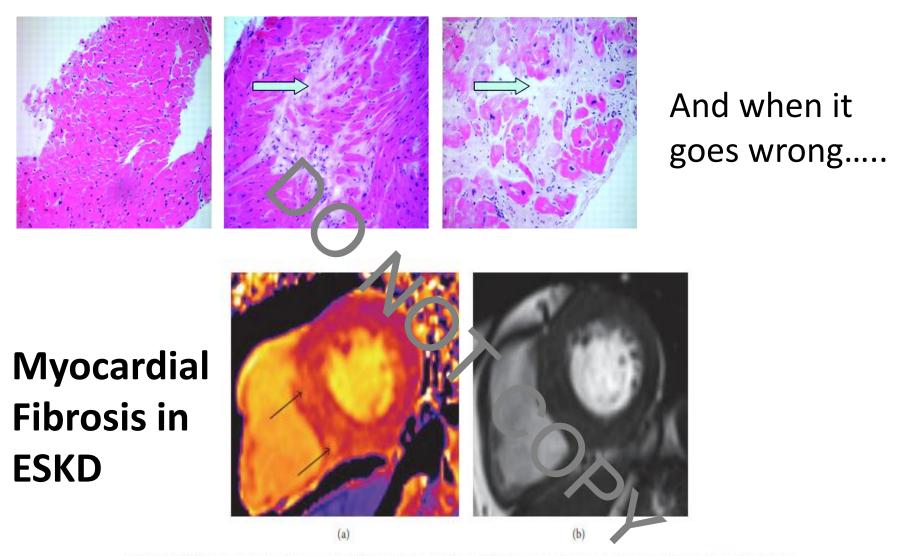
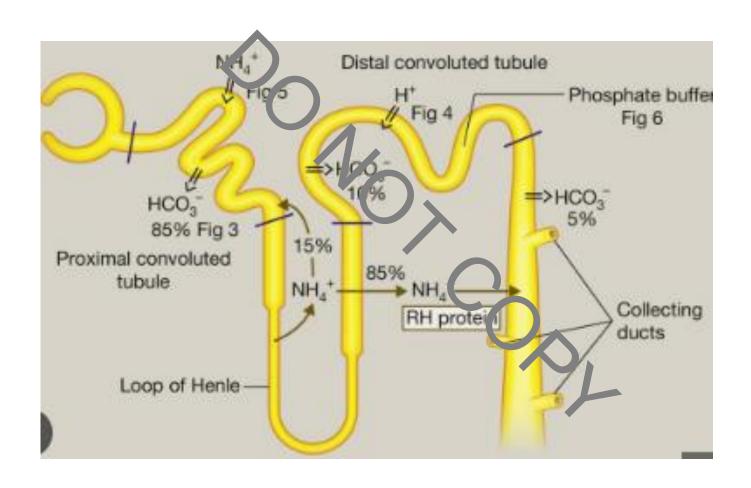


FIGURE 4: (a) Short-axis midventricular native T1 map of a dialysis patient. Black arrows show areas of discretely increased signal intensity likely to represent myocardial fibrosis. (b) Corresponding short-axis midventricular plain cardiac MRI cine image of the left ventricle of the same dialysis patient. No tissue abnormality visible on plain MR imaging.

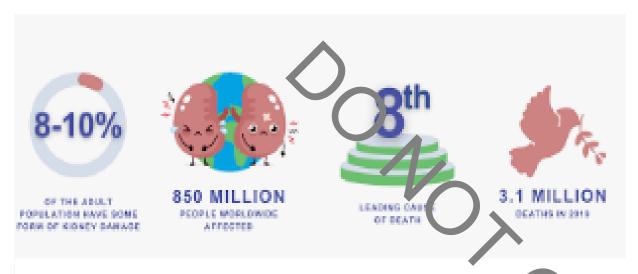
Graham-Brown MPM et al, Biomed Res Int 2017

Functions of the kidneys



Crucial in acid-base balance

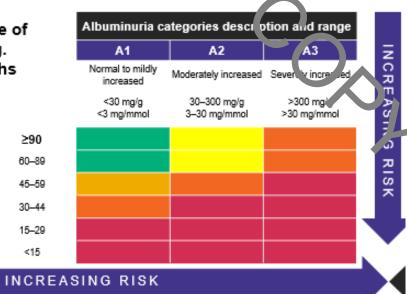
Functions of the kidneys



Glomerular filtration of waste

CKD is defined as GFR <60 ml/min/1.73 m² or presence of markers of kidney damage, e.g. ACR ≥3 mg/mmol, for >3 months

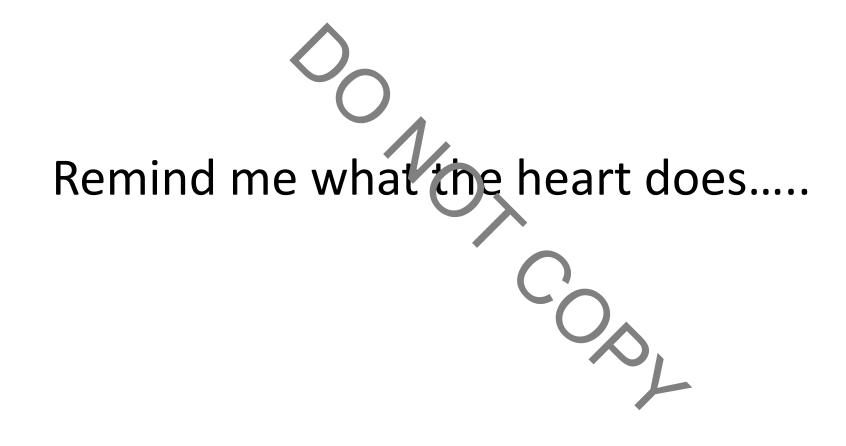
GFR categories (ml/min/1.73 m²) description and range	G1	Normal or high	≥90
	G2	Mild	60-89
	G3a	Mild to moderately decreased Moderately to severely decreased	45–59
	G3b		30-44
	G4	Severely decreased	15–29
	G5	Kidney failure	<15

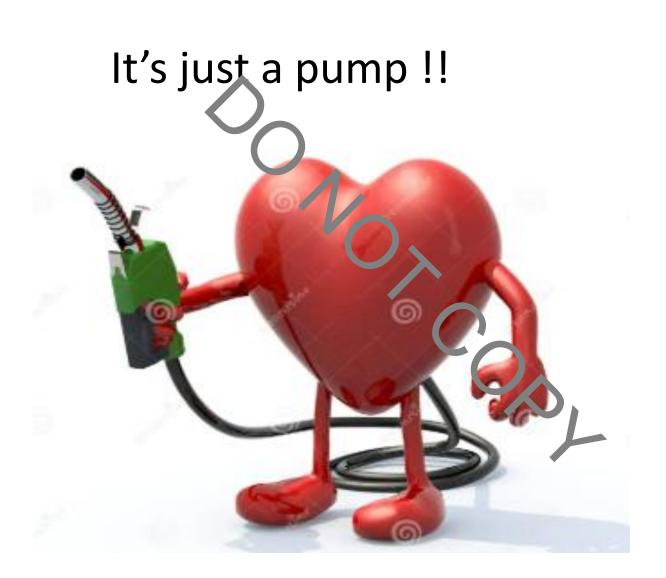


Functions of the kidney: Water balance

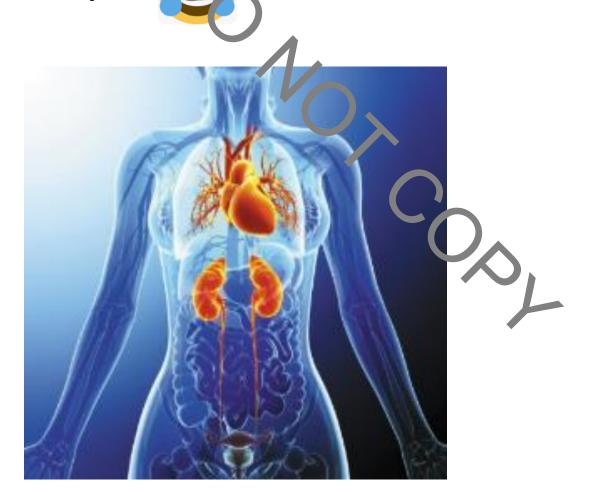
Where would heart failure management be without a decent diuresis!!



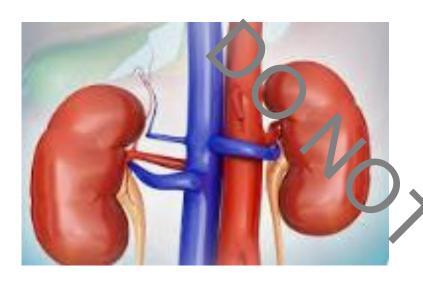




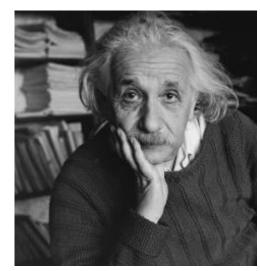
But to be fair – that pump does feed the kidneys



There's no competition!!



They really are the smartest



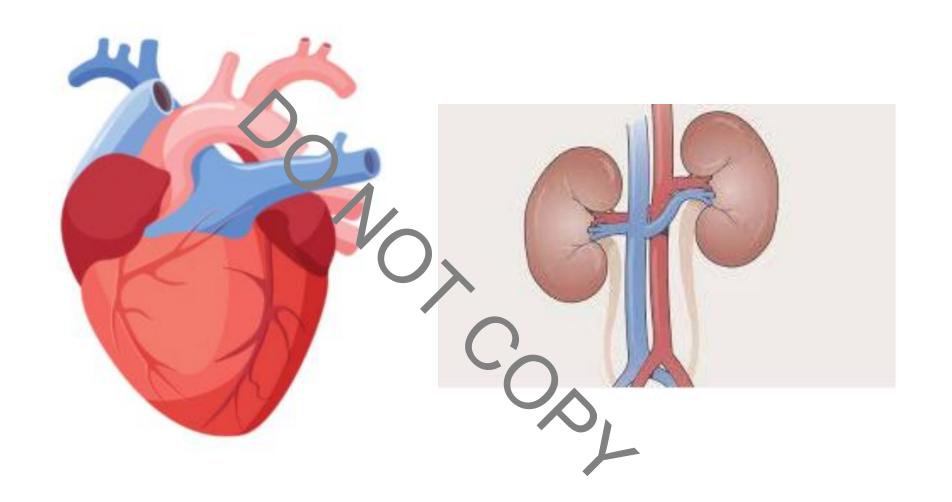


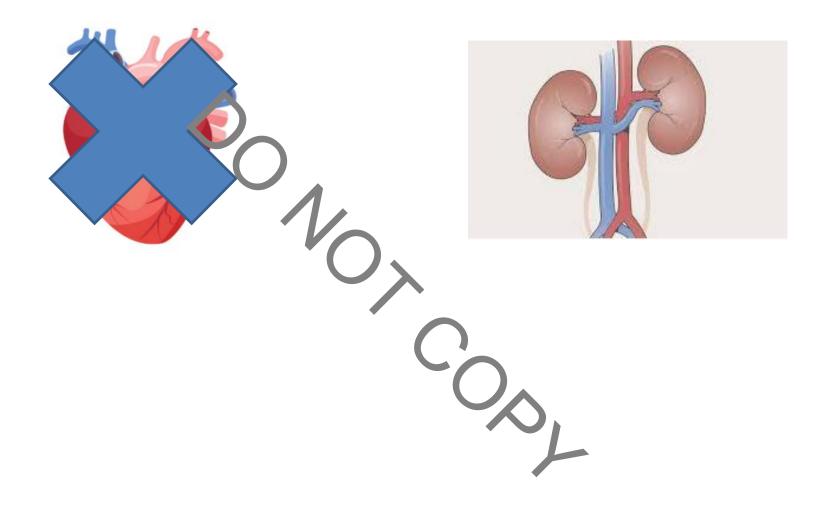
What is more important, the kidneys or the heart?

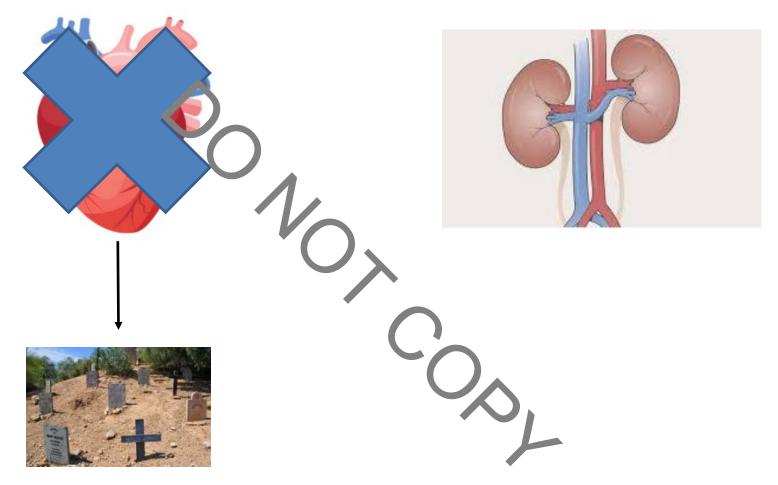
Paui R Kalra

Consultant Cardiologist

Portsmouth Hospitals University NHS Trust

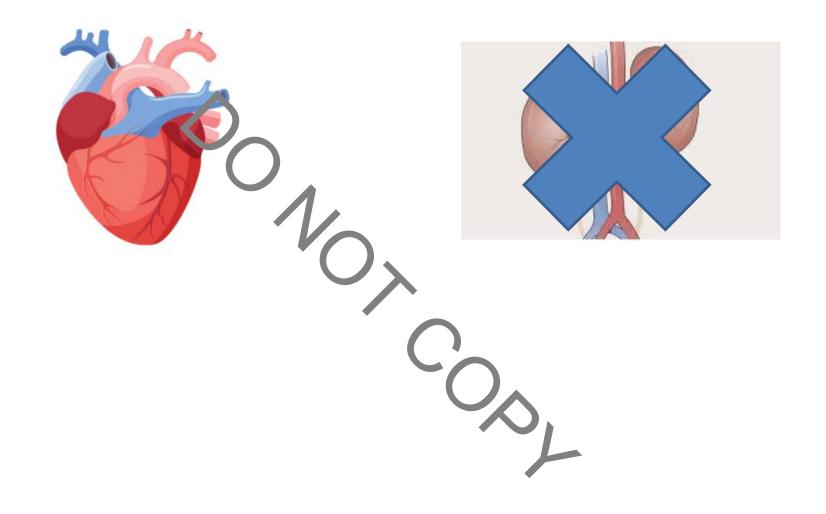


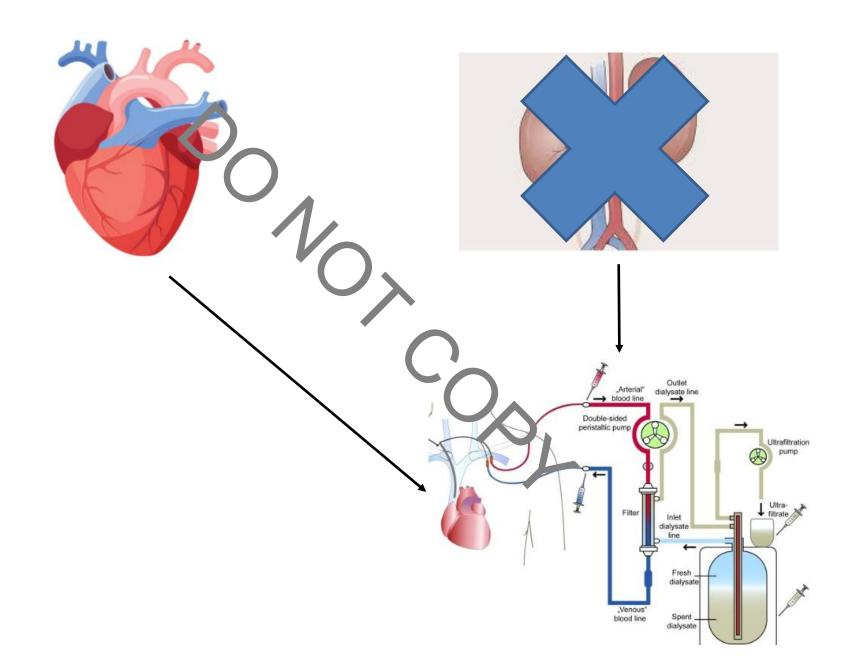




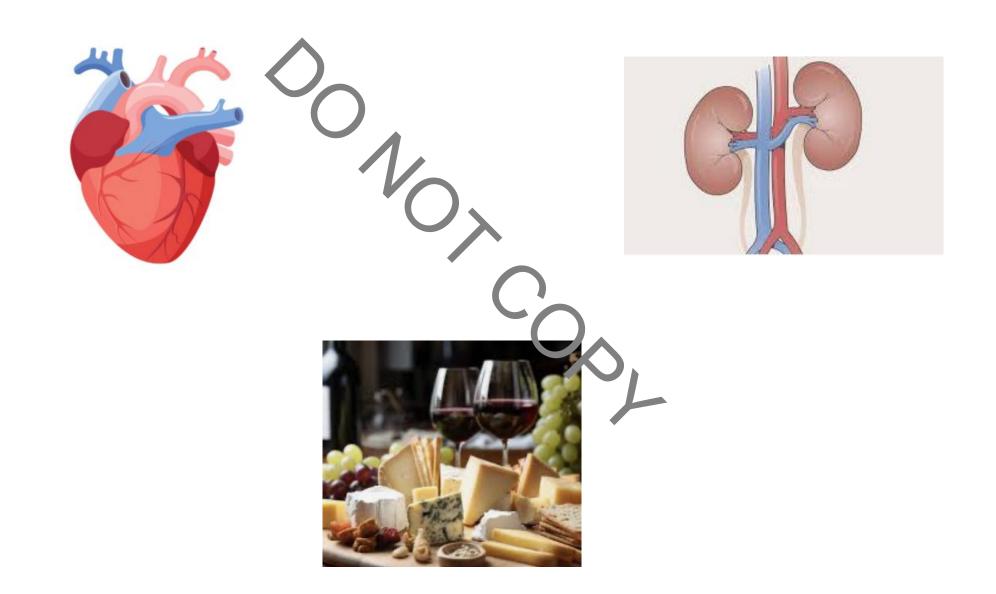








Options to vote now on what is most important...



• 75-year-old man, HFrEF

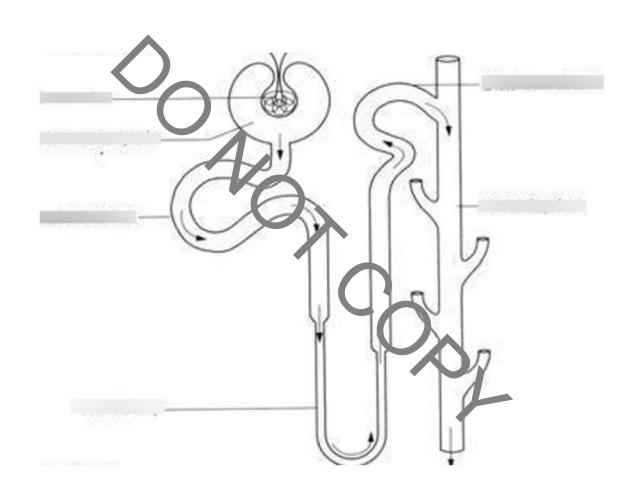
LRTI 2 weeks before and then progressive dyspnoea and leg swelling

Usual eGFR 40-45, currently 32

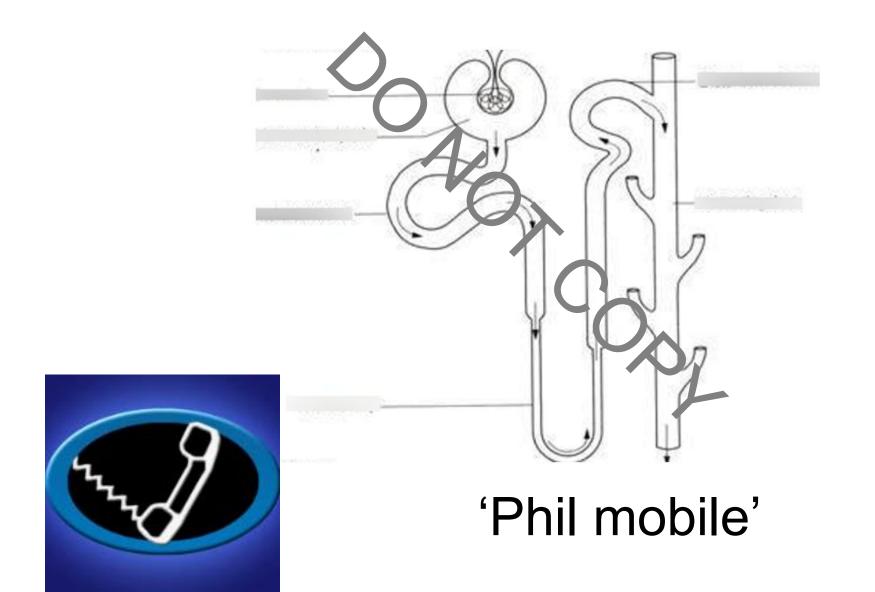




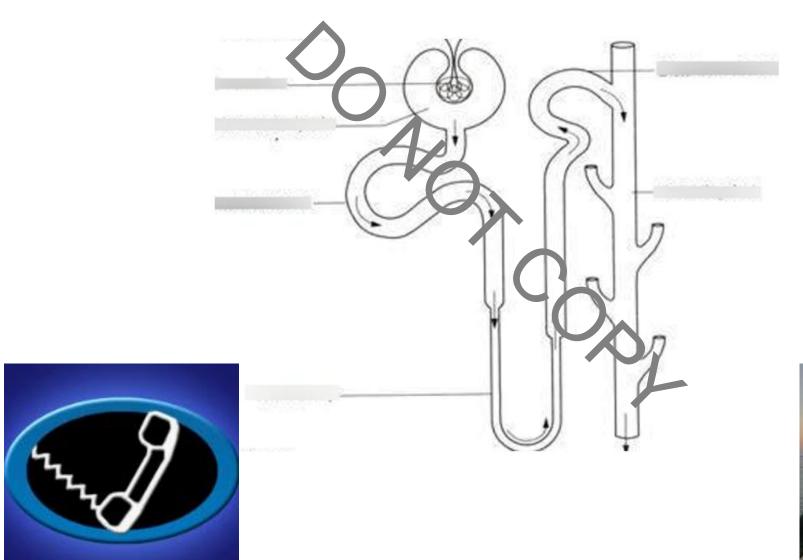
Nephron anatomy and blockade



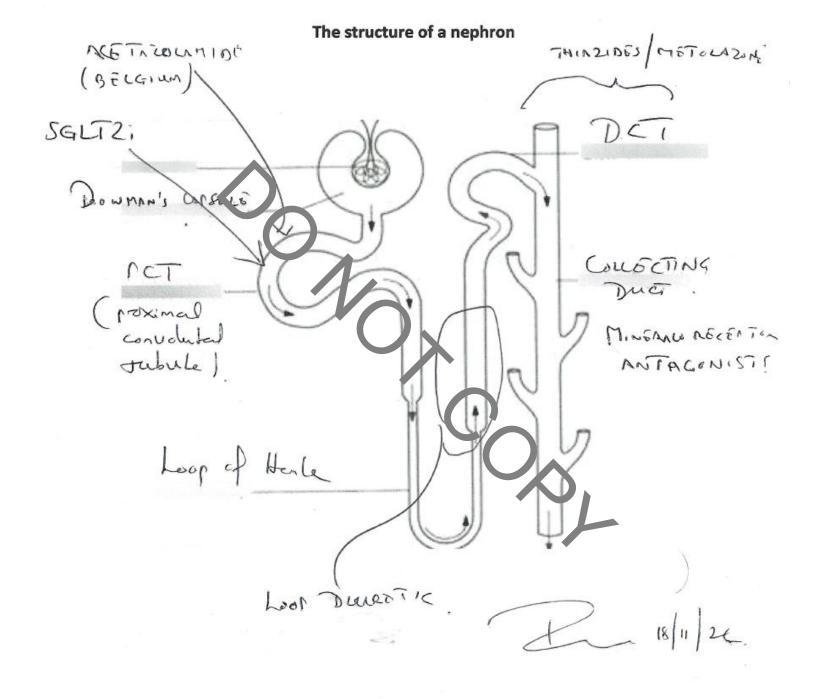
Nephron anatomy and blockade



Nephron function and blockade







Case study: Patient presentation



IBRAHIM, 69-YEAR-OLD MALE SHOP OWNER

Weight 93 kg / BMI 34 kg/m2



Blood pressure

- In practice: 158/88 mmHg
- Home monitoring: average 152/90 mmHg from recent readings
- Haemoglobin : 108 g/L



ECG: sinus rhythm, 78/min; normal QRS **NT pro-BNP** 675 ng/l

Medical history

- Hypertension and type 2 diabetes for 5 years
- Transient ischaemic attack 2 years earlier
- No regular exercise but working >60 hours/week
- Last reviewed 2 years earlier

Current medication

- Metformin 1000 mg BD
- Linagliptin 5 mg OD
- Gliclazide 80 mg BD
- Irbesartan 150 mg OD
- Amlodipine 10 mg OD
- Atorvastatin 10 mg OD



Lipid profile

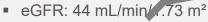
- Total-sholesterol: 5.8 mmol/L
- LDL: 3.3 mmol/L
- HDL: 1.2 mmol/L
- Non-HoL: 4. mmol/L
- Triglyc rider. 3/2 mmol/L



Liver function

- LFTs:
 - ALT: 42 U/L
 - Alk phos: 97 U/LBilirubin: 12 umol/l





- Sodium: 136 mmol/L
- Potassium: 5.2 mmol/L
- Creatinine: 140 µmol/L
- Urine ACR: 68 mg/mmol



Glucose

HbA1c: 71 mmol/mol

Current presentation

 Presents complaining of breathlessness on walking uphill and upstairs and some ankle oedema

ACR, albumin:creatinine ratio; ALT, alanine transaminase; AST, aspartate aminotransferase; BD, twice daily; eGFR, estimated glomerular filtration rate; HbA1c, glycated haemoglobin A1c; HDL, high-density lipoprotein; LDL, low-density lipoprotein; LFT, liver function test; OD, once daily; QRISK, Cardiovascular Risk Score; U+E, urea and electrolyte

Cardio-renal-metabolic management

- Renal USS: both kidneys 10.3 cm; decreased cortico-medulary differentiation; bladder emptied fully
- Serum free light chains normal
- Ferritin 67 ug/l and TSat17%
- Echocardiogram : LVEF 40% with global hypokinesia

Cardio-renal-metabolic management

Investigations

- Renal USS: both kidneys ~ 10.3 cm; decreased cortico-medulary differentiation; bladder emptied fully
- Serum free light chains normal
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Key management

- Kidneys: BP control; decrease albuminuria; slow CKD progression
- Heart: 4 pillars of HF management
- CV risk: LDL control, weight loss and exercise
- Metabolic : glycaemic control and weight loss

Evidence based Cardio-renal-metabolic management

Kidney management

- Target BP of ~ 130/80
- maximise Renin angiotensin blockade(RAB)
- irbesartan 300 mg OD (IDNT)
- <u>+</u> finerenone (Fidelio)
- SGLT2i eg dapagliflozin (DAPA-CKD) or empagliflozin (EMPA Kidney)

Heart failure management

- 4 pillars of HF Rx : RAB; ß blocker; MRA; SGLTi (DAPA-HF and EMPA-Reduced)
- Sacubitril valsartan vs irbesartan (PARADIGM-HF)
- IV iron (IRONMAN)

CV risk management

- Control LDL cholesterol to < 1.8 mmol/l (atorvastatin 40-80 mg OD)
- GLP1 agonist eg semaglutide sc (FLOW)
 - Exercise and weight loss

Metabolic management

- SGL721
- GLP1 agenist
- Exercise and weight loss
- Aim for HbA1C < 53 mmol/mol

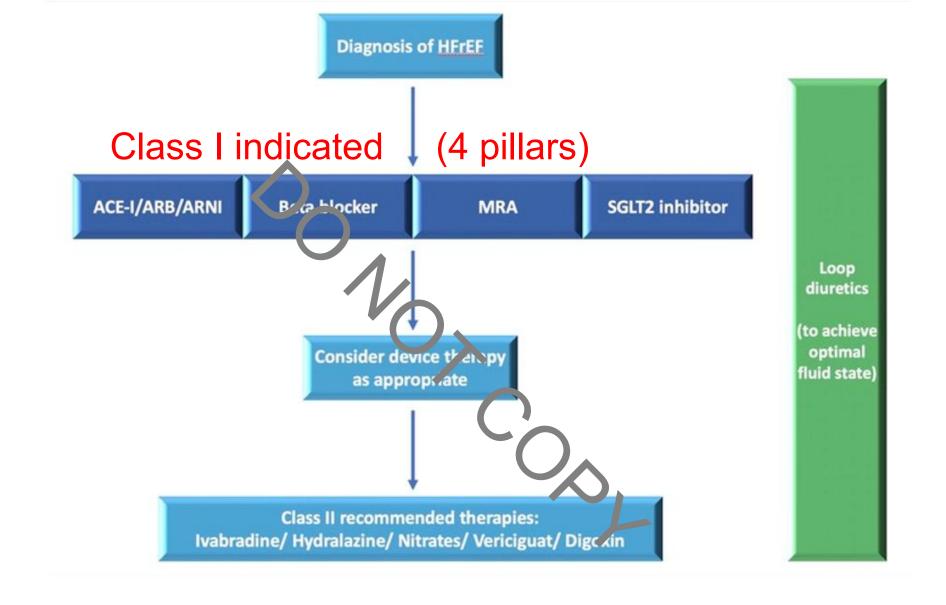


Figure 1. Adapted from 2021 ESC HF guidelines. Algorithm for the management of HFrEF.

Adapted from: McDonagh et at. Eur Heart J 2021;42(36):3599-3726.

• 75-year-old man, HFrEF

LRTI 2 weeks before and then progressive dyspnoea and leg swelling

Usual eGFR 40-45, currently 32





75-year-old man, HFrEF

LRTI 2 weeks before and then progressive dyspnoea and leg swelling

Usual eGFR 40-45, currently 32

Diagnosis: AKI



75-year-old man, HFrEF

 LRTI 2 weeks before and then progressive dyspnoea and leg swelling

Usual eGFR 40-45, currently 32

Diagnosis: <u>high risk heart failure with AKI</u>

CKD is an independent predictor of adverse outcome in CHF

CHARM: Adjusted rates of CV death or hospitalisation for CHF (n=2680)

eGFR (mL/min/1.73m²)		lazard ratio	95% CI	Р
≥90	(n=577)	1.00	•••	•••
75–89.9	(n=519)	1.17	0.93–1.49	0.180
60–74.9	(n=618)	1.24	0.98–1.56	0.070
45–59.9	(n=547)	1.54	1.22–1.94	<0.001
<45	(n=419)	1.86	1.47-2.36	<0.001

Decongestion, decongestion

Congestion is awful for patients and for the kidneys

 Need much higher dose than usual dose (irrespective of if oral or IV)

Low dose if renal function QK and diuretic naïve

 If no improvement, then step up therapy (combination of loop, SGLT2i, MRA and thiazide)

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Consider renal USS and urine ACR

FACTS

Angiotensin converting enzyme inhibitors, angiotensin receptor blockers, mineralocorticoid receptor antagonists, SGLT2 inhibitors are NOT nephrotoxins

They are indeed nephroprotective drugs (that affect renal blood flow)

Change in renal function associated with drug treatment in heart failure: national guidance

Andrew L Clark, Paul R Kalra, Mark C Petrie, Patrick B Mark, Laurie A Tominson, Charles RV Tomson

Consensus statement from the British Society for Heart Failure and the Renal Association





Heart 2019;**105**:904–910.

Management of renal function during initiation of RAAS inhibitors

Table 1 Management of RAAS inhibitors is response to change in renal function

Clinical assessment:

- Compare with baseline renal function (review series of results).
- Assess fluid status: if intravascularly depleted (jugular venezus alse not visible, postural drop in BP and no oedema), consider cautious intravenous fluids.
- ► Interpret BP in the context of usual values (low BP does not necessar of nean patient needs fluid).
- Reduce/withdraw RAASI if symptomatic hypotension.
- Repeated clinical and biochemical assessment is vital.
- Presence of moderate or severe hyperkalaemia may override recommendations based on change in renal function.
- In severe renal dysfunction assess for symptoms or uraemia.

	Recommendations for kAAS inhibitors		
Change in renal function compared with baseline	HFpEF (assuming no other prognostic i) dication).	HFREF.	
Increase in serum creatinine by <30%	Consider stop ACEI/ARB/ARNI Review MRA according to fluid status.	Continue unless symptomatic hypotension.	
Increase in serum creatinine 30%-50%	Stop RAAS inhibitor.	Consider reducing dose or temporary withdrawal.*	
Increase in serum creatinine >50%	Stop RAAS inhibitor.	Temporarily stop RAAS inhibitor.*	
Severe renal dysfunction, for example, eGFR <20	Stop RAAS inhibitor.	top RAAS inhibitor if symptomatic uraemia irrespective of baseline function.	

^{*}Reinitiate and/or retitrate when renal function improved in patients with HFrEF

What is more important : the Kidneys or the Heart?



What is more important: the Kidneys or the Heart?

