

CRRT Concetti di base 3

Stefano Romagnoli, MD, PhD



Dip. di Scienze della Salute – Università di Firenze Dip. di Anestesia e Rianimazione - AOU Careggi - Firenze **CRRT: concetti di base 1**: come funzionano le CRRT. Come funziona il circuito extracorporeo? Quali sono i meccanismi di trasporto soluti e acqua nei filtri? | *Z. Ricci*



CRRT: concetti di base 2: Nomenclatura: descrizione delle varie componenti del circuito, membrane, filtri, parti delle apparecchiature e sensori | *M. Neri*

Agenda

• <u>Tecniche</u>:

- descrizione delle varie tecniche e delle loro caratteristiche sulla base di:
 - Efficienza
 - Peculiarità
 - Impiego clinico
 - Modalità applicative



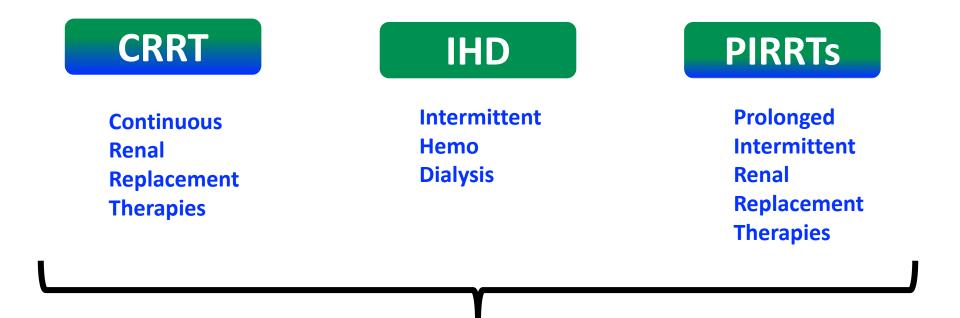
Continuous Renal Replacement Therapy Who, When, Why, and How ≋CHEST

Modalities of RRT

Multiple modalities of renal support may be used in the management of the **critically ill patient** with kidney failure.



Tandukar S & Palewsky PM. CHEST 2019;155:626-638



All of these use relatively similar extracorporeal blood circuits and differ primarily with regard to <u>duration of therapy</u> and, consequently, the **rapidity of net ultrafiltration and solute clearance**.

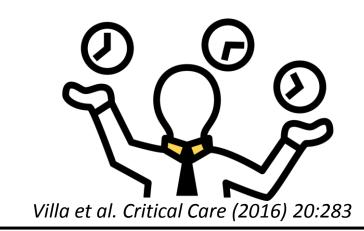
Tandukar S & Palewsky PM. CHEST 2019;155:626-638

Hybrid therapies - PIRRT

 With respect to frequency and duration, the term "hybrid therapies" relates to the <u>blending or characteristics from both intermittent and continuous</u> <u>modalities</u>.

- These therapies attempt to <u>optimize the advantages and minimize the</u> <u>disadvantages</u> of both modalities:
 - <u>Efficient solute removal</u>
 - Slower ultrafiltration rates for hemodynamic stability
 - Less anticoagulant exposure
 - Shorter duration
 - Lower costs
 - Decreased nurse workload

Nomenclature for renal replacement therapy and blood purification techniques in critically ill patients: practical applications



Nomenclature for RRT: practical applications

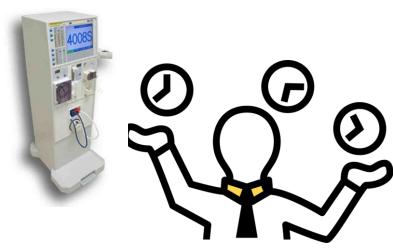
Hybrid therapies - PIRRT

Hybrid therapies encompass various specific "discontinuous" RRT modalities: :

- ✓ Sustained low-efficiency dialysis (SLED),
- Slow low-efficiency extended daily dialysis (SLEDD),
- Extended daily dialysis (EDD),
- Extended daily dialysis with filtration (EDDf),
- Extended dialysis (ED),
- ✓ "go slow dialysis",
- ✓ Accelerated veno-venous hemofiltration (AVVH).

Hybrid therapies are **usually performed with** standard intermittent hemodialysis equipment (machines, filters, extracorporeal blood circuits). Solute removal is largely diffusive but variants with a convective component, such as EDDf and AVVH, are possible.

Nomenclature for renal replacement therapy and blood purification techniques in critically ill patients: practical applications



Villa et al. Critical Care (2016) 20:283 (modified)

Nomenclature for RRT: practical applications

Prolonged Intermittent Renal Replacement Therapy

	INTERMITTENT	CONTINUOUS	HYBRID
	IHD	CRRT	PIRRT
Solute clearance	Diffusion	Diffusion / Convection / Both	Diffusion / Convection / Both (Most PIRRT trials have reported a diffusive modality)
Type of machine	Standard IHD machine	Standard CRRT machine	Either IHD or CRRT machine (Most PIRRT trials have used an IHD machine).
Qb (ml/min)	400-500	100-200	150-400
Qd (ml/min)	600-800	(CVVHD – CVVH – CVVHDF)	100-200
Duration (h)	3-4	24	6-12

Edrees F et al. Advances in Chronic Kidney Disease; 23;3, 2016:195-202(modified)

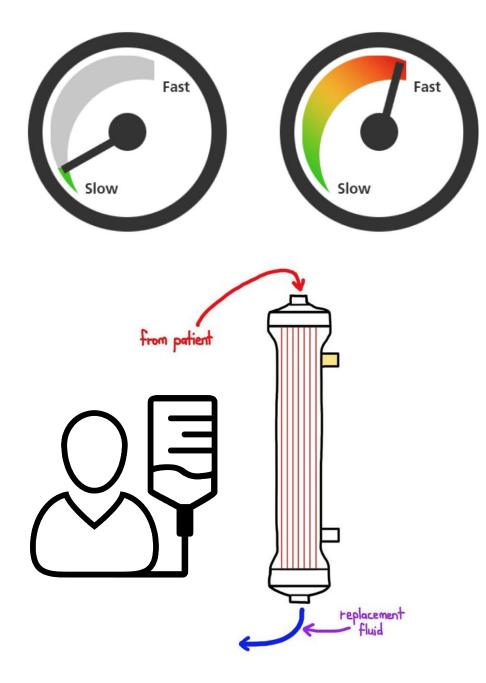
Prolonged Intermittent Renal Replacement Therapy

Prolonged Intermittent Renal Replacement Therapy

	INTERMITTENT	CONTINUOUS	HYBRID
	IHD	CRRT	PIRRT
Frequency	2-3 d/wk	Continuous	3-7 d/wk
Timing	Day	Day + night	Day or night
Anticoagulation	Can be performed without anticoagulation	Usually yes	Can be performed without anticoagulation
Vascular access	AVF – AVG - CVC	CVC	CVC
Usual UF rate	1-5 L/3-4 h	1800-2500 ml/h	1-4 L/6-12 h
Patient location	ICU, step down unit, ward	ICU, step down unit	ICU, step down unit, ward

Edrees F et al. Advances in Chronic Kidney Disease; 23;3, 2016:195-202(modified)

Prolonged Intermittent Renal Replacement Therapy





IHD provides **rapid** solute **clearance** and **ultrafiltration** during relatively brief (3- to 5-h) treatments

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WHAT IS THE OPTIMAL RRT MODALITY FOR PATIENTS WITH AKI?

- The most problematic of these is intradialytic hypotension, which results from fluid removal on dialysis <u>at a rate that exceeds</u>
 <u>vascular refilling</u> from the intracellular and interstitial compartments.
- Intradialytic hypotension is a common complication in critically ill patients who often have the vexing combination of fluid overload and hemodynamic instability.

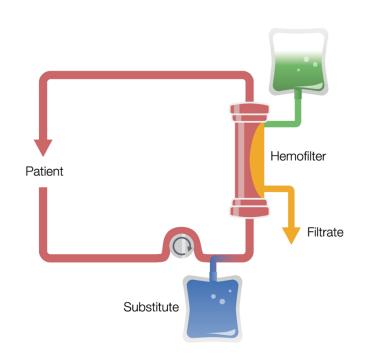


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WHAT IS THE OPTIMAL RRT MODALITY FOR PATIENTS WITH AKI?

CRRT seems like an ideal modality for dialysis of critically ill patients, as it would be predicted to cause—by virtue of its slow, continuous nature—less hypotension and less radical electrolyte and pH perturbation than IHD.

Guérin C et al. Intensive Care Med 2002; 28:1411–1418 Bellomo R et al. Nephron 1995; 71:59–64

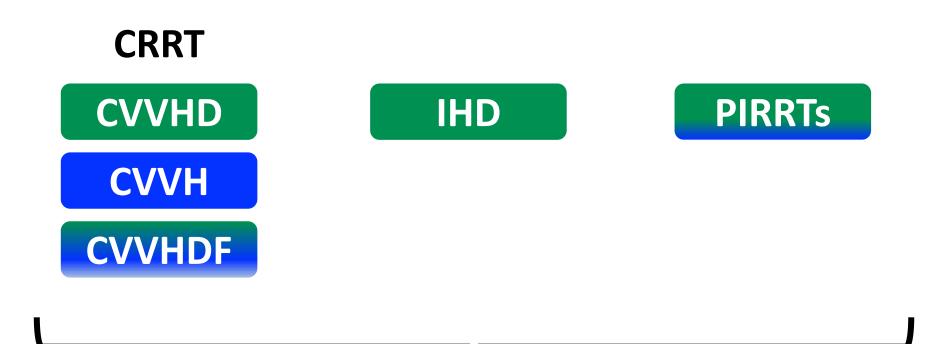


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The continuous therapies provide **more gradual fluid removal** and **solute clearance** over <u>prolonged treatment times</u> (optimally, 24 h per day but often interrupted due to system clotting or diagnostic or therapeutic procedures)

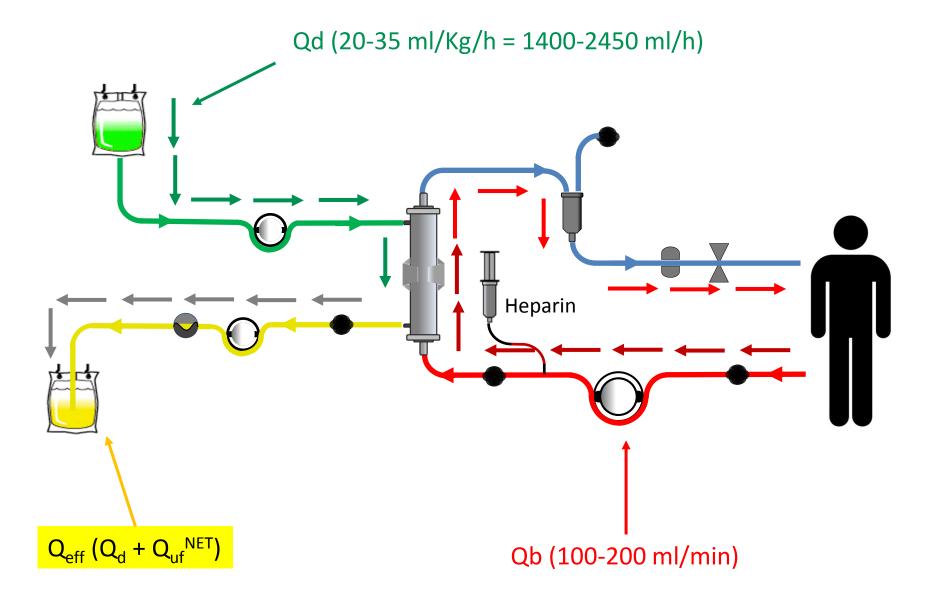
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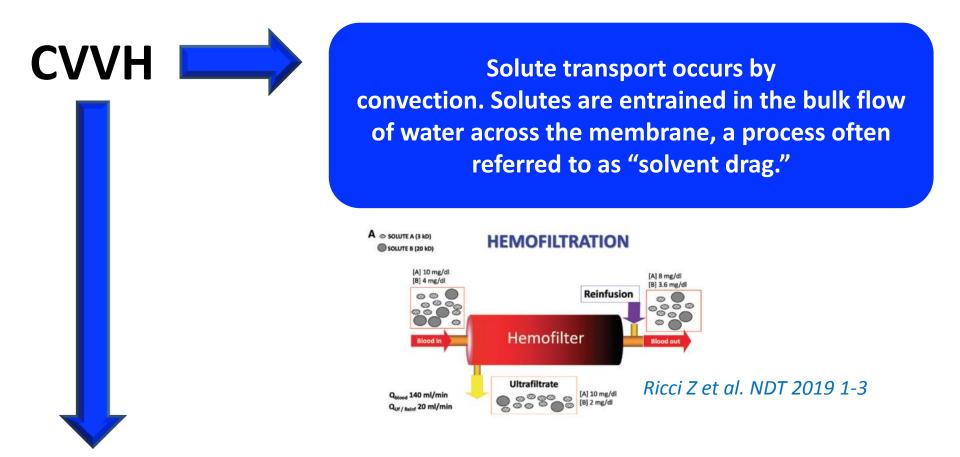


In addition, dialytic therapies rely predominantly on **diffusive** solute clearance, whereas solute removal during hemofiltration occurs by **convection**.

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CVVHD heparin

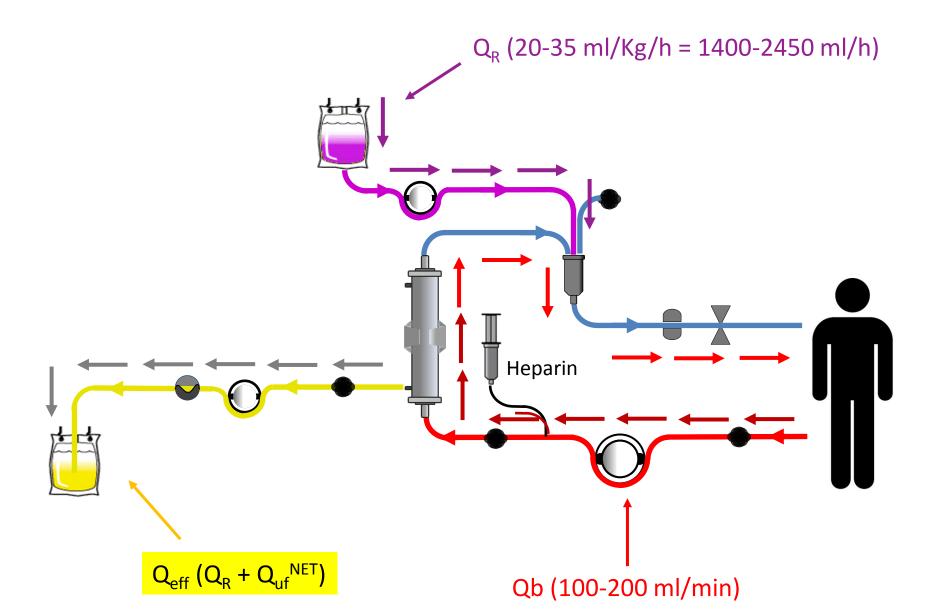




High ultrafiltration rates are needed to achieve sufficient solute clearance, and the ultrafiltrate volume beyond what is required to achieve desired net fluid removal is replaced with balanced IV crystalloid solutions.

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CVVH (100% post) heparin



CVVH (Continous Veno-Venous Hemofiltration)

- Continuous veno-venous hemofiltration (CVVH) uses convection to remove solutes through large volume fluid ultrafiltration.
- Small solute molecules, such as urea, <u>and middle-sized</u> <u>molecules</u>, such as inflammatory **cytokines**, are cleared.
- With the large volume of fluid removed, intravascular volume must be maintained using a **replacement fluid**.

Alvarez G et al. Can J Anesth 2019 Feb 6.

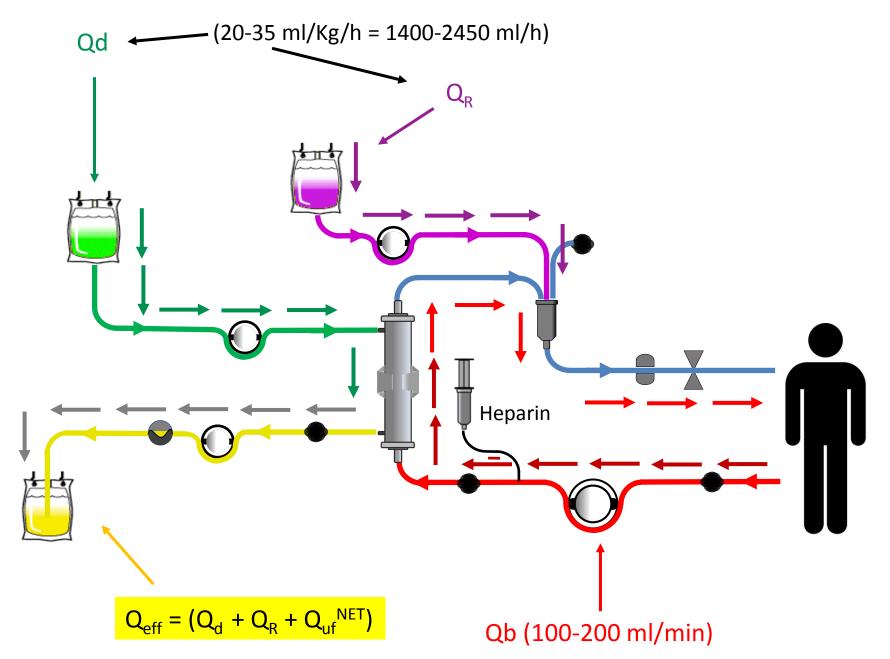
Review - overview CRRT

CVVH (Continuous Veno-Venous Hemofiltration)

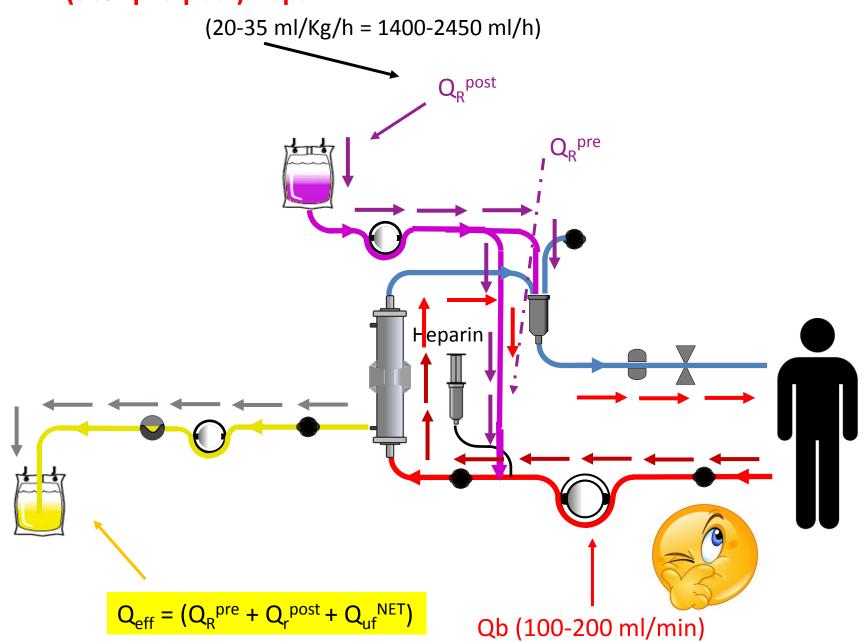
- The replacement fluid can be infused either before the hemofilter (predilution) or after the hemofilter (postdilution).
- Postdilution results in more concentrated blood in the filter and higher solute clearance.
- Nevertheless, more concentrated blood can lead to a shorter filter lifespan.
- While pre-dilution means lower solute concentrations and clearance, this is offset by a higher ultrafiltration rate and **longer filter life**.

Alvarez G et al. Can J Anesth 2019 Feb 6.

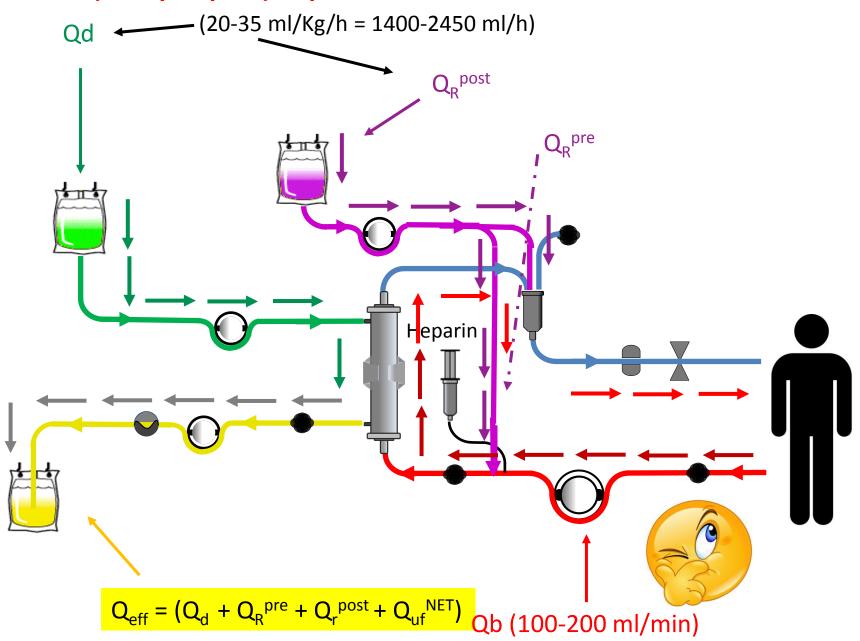
CVVHDF (100% post) heparin



CVVH (50% pre-post) heparin



CVVHDF (50% pre-post) heparin



Acute kidney injury: to dialyse or to filter?

Zaccaria Ricci^{1,*}, Stefano Romagnoli² and Claudio Ronco^{3,4}



<u>As a practical approach</u>, in order to achieve <u>advantages from both</u> <u>techniques (Diffusion & Convection)</u>:

- Set: CVVHDF
- Prescription 20 35mL/kg/h of dialytic dose:
 - **Split** the flows between **haemofiltration** and **dialysis** in order to <u>balance</u> prolonged <u>circuit life</u> and <u>efficiency</u> of blood purification.

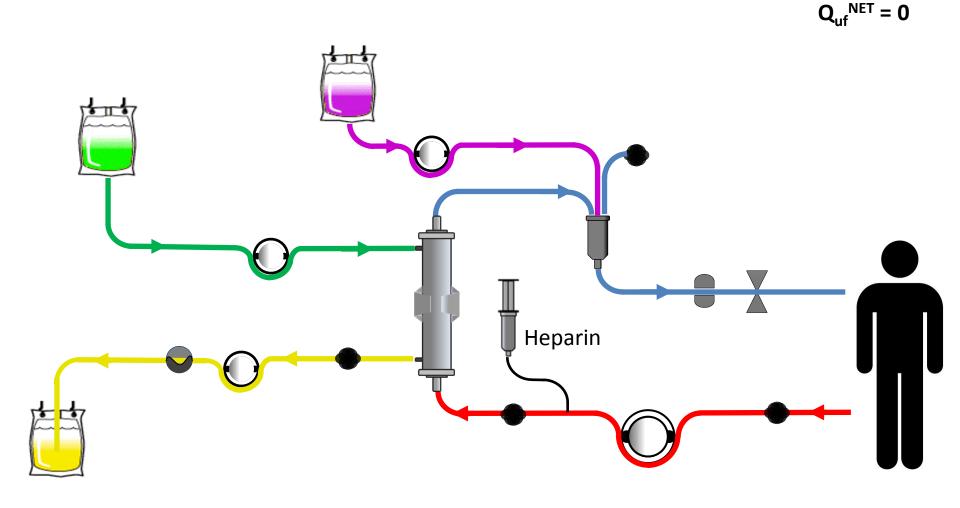
However, during haemodiafiltration, the **highest possible convective dose should always be delivered by setting the** <u>haemofiltration rate</u> <u>to 20% of plasma flow</u>, with the remaining prescription set as diffusion, just to reach the desired intensity target.

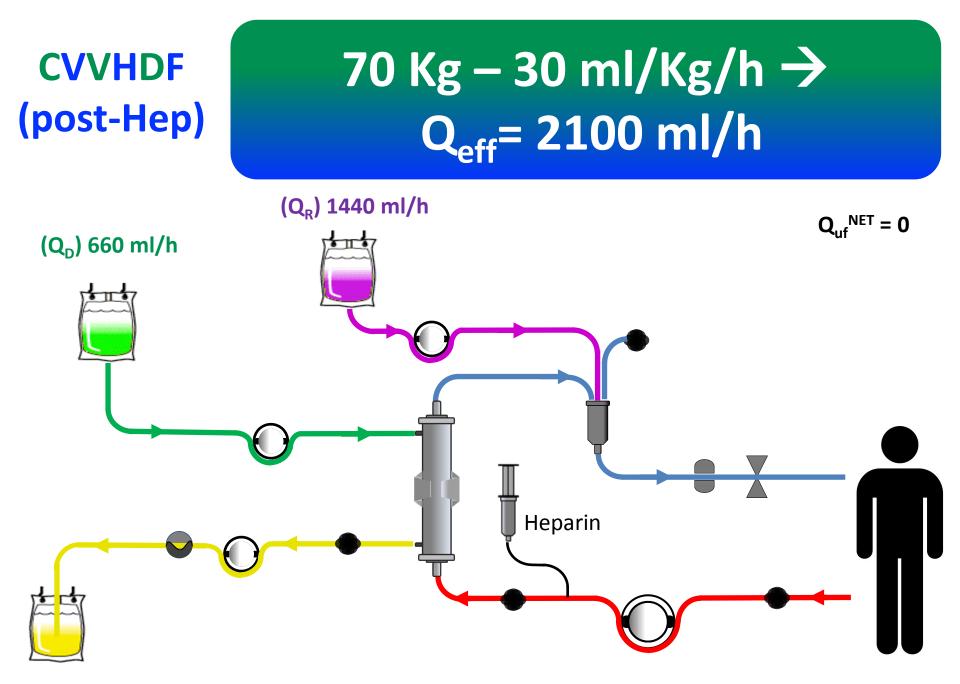
Ricci Z et al. Nephrol Dial Transplant (2019) 1–3

Overview exploring Diffusion and Convection

CVVHDF (post-Hep)

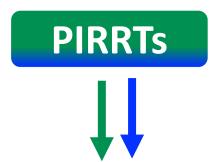
70 Kg – 30 ml/Kg/h \rightarrow Q_{eff}= 2100 ml/h





 $Q_b = 200 \text{ ml/min} \rightarrow Qp = 120 \text{ ml/min} \rightarrow 7.200 \text{ ml/h} \rightarrow (20\%) \rightarrow Quf = 1440 \text{ ml/h}$





The multiple forms of PIRRT are characterized by treatments that are generally **between 8 and 16 h in duration**, with **slower** rates of solute <u>clearance and ultrafiltration</u> <u>than IHD</u> but <u>more rapid than</u> <u>CRRT</u>.

Tandukar S & Palewsky PM. CHEST 2019;155:626-638

PIRRT is most commonly provided by using **equipment similar to that for IHD** but with lower blood and dialysate flow rates.



It can also be performed by using **equipment designed for CRRT** but with **augmented dialysate and/or ultrafiltration rates** to achieve <u>similar delivered therapy over a shorter duration</u>



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WHAT IS THE OPTIMAL RRT MODALITY FOR PATIENTS WITH AKI?



 There are no convincing data that demonstrate a difference in important clinical outcomes with PIRRT, compared with other RRT modalities

Edrees F et al. Adv Chronic Kidney Dis. 2016;23(3):195-202.



Rachoin JB & Weisberg LS. CCM 2019 Feb 13

Selection of RRT Modality

Although **CRRT** and **PIRRT** are most commonly used in **hemodynamically unstable patients**, there is marked variation in practice



Some centers use CRRT (or PIRRT) in all ICU patients with renal failure regardless of hemodynamic status, whereas **others** use IHD, albeit with adjustments in prescription, even in vasopressor dependent patients.

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Although the benefit of a slow, continuous modality of renal support in hemodynamically unstable patients may seem **Self-evident**, randomized trials have failed to show differences with regard to either mortality or recovery of kidney function comparing CRRT with either IHD or PIRRT

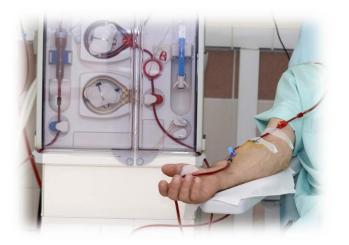
Mehta RL et al. Kidney Int. 2001;60(3):1154-1163. Augustine JJ, et al. Am J Kidney Dis. 2004;44(6):1000-1007. Uehlinger DE, et al. Nephrol Dial Transplant. 2005;20(8):1630-1637. Vinsonneau C et al. Lancet. 2006;368(9533):379-385. Lins RL et al. Nephrol Dial Transplant. 2009;24(2):512-518. Schefold JC et al. Crit Care. 2014;18(1):R11. Bagshaw SM et al. Crit Care Med. 2008;36(2):610-617. Pannu N et al. JAMA. 2008;299(7):793-805. Friedrich JO et al. Critical Care. 2012;16(4): R146. Zhang L et al. Am J Kidney Dis. 2015;66(2):322-330. Kielstein JT, et al. Am J Kidney Dis. 2004;43(2):342-349. Schwenger V et al. Crit Care. 2012;16(4):R140. KDIGO. Kidney Int. 2012;2012(suppl):1-138.

It <u>must be recognized</u>, however, that to provide **IHD** in hemodynamically unstable patients, the standard prescription may require modification, such as <u>prolongation of treatment time</u> to allow for more gradual ultrafiltration, use of higher dialysate sodium concentrations, and reduced dialysate temperatures

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WHAT IS THE OPTIMAL RRT MODALITY FOR PATIENTS WITH AKI?

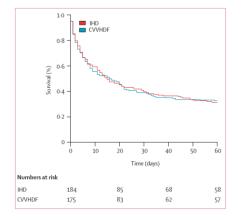
In order to minimize hemodynamic perturbation in critically ill patients on IHD, the prescription may need to be modified.



 Exacerbation of hemodynamic instability may be mitigated by extending treatment time

[Hemodiafe trial: average lenght = 5.2 hr per session]

Vissonneau C et al. Lancet. 2006 29; 368(9533): 379-85



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Although the Kidney Disease: Improving Global Outcomes (KDIGO) Clinical Practice Guideline for AKI recommends the use of CRRT for patients who are hemodynamically unstable, the strength of this recommendation is low.

Observational data, however, do suggest that CRRT is more effective in **achieving net** <u>negative fluid</u> <u>balance</u> than IHD.



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The various mechanisms of solute clearance provided by CVVH and CVVHD result in different profiles of solute removal with each modality.

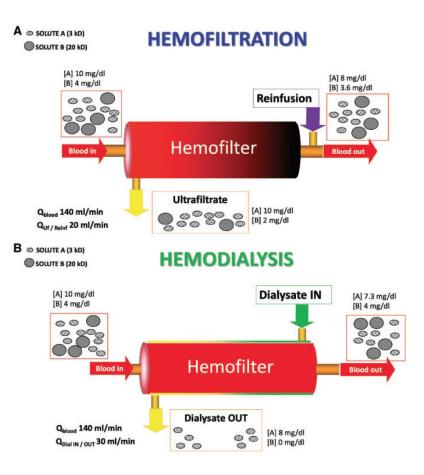
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Nephrol Dial Transplant (2019) 1–3 doi: 10.1093/ndt/gfz022

Acute kidney injury: to dialyse or to filter?

Zaccaria Ricci^{1,*}, Stefano Romagnoli² and Claudio Ronco^{3,4}

Extraction of mediumsized and large molecules from the blood is greater with convective rather than diffusive methods.



Ricci Z, Romagnoli S, Ronco C. NDT 2019 1-3

Review on Solute Transport

CVVHD IHD

Diffusion provides efficient clearance of lowmolecular-weight solutes (< 500-1,500 Daltons)

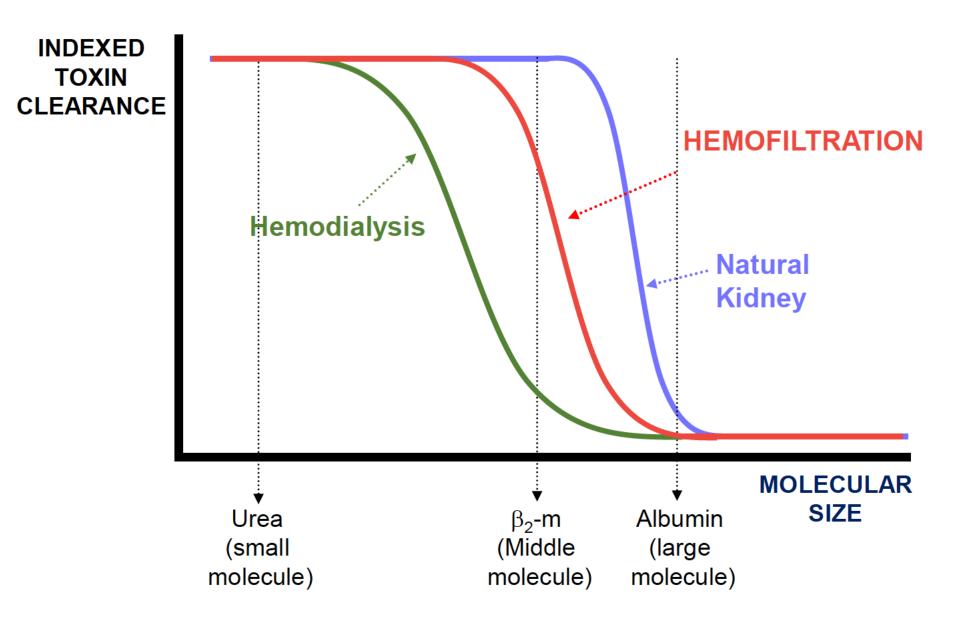
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CVVH

Solute movement in convection is limited primarily by the size of the **pores** in the hemofilter membrane.

CVVHDF

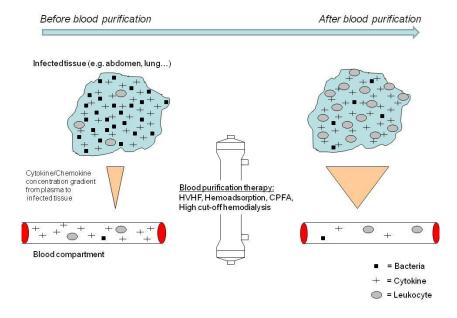
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WHAT IS THE OPTIMAL RRT MODALITY FOR PATIENTS WITH AKI?

 Several studies have examined effects of convective modalities on cytokines in patients with sepsis and AKI

Bellomo R et al. Ren Fail 1995; 17:457–466 Kellum JA et al. Crit Care Med 1998;26:1995–2000 Lonnemann G et al. Kidney Int Suppl 1998; 66:S43–S46 van Bommel EF et al. Ren Fail 1997; 19:443–454

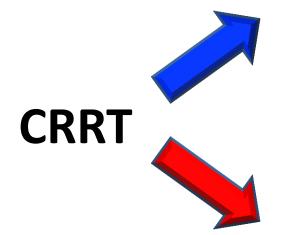


Rimmeleé T & Kellum JA. Critical Care201115:205

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CRRT in Sepsis and Multisystem Organ Failure

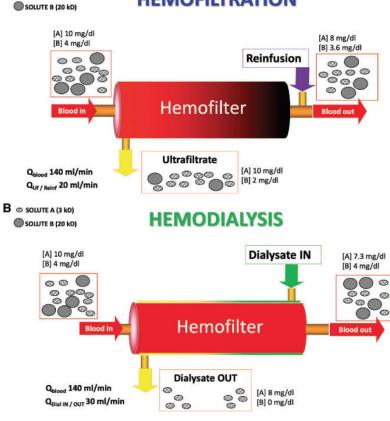
Providing more time to achieve fluid balance and metabolic homeostasis



Modulator of the immune response.

- High-volume hemofiltration (HVHF)
- High-cutoff membranes
- Hybrid (filtration + adsorption) systems such as coupled plasma filtration absorption.

Macedo E et al. Am J Kidney Dis. 2016;68:645-65



Ricci Z, Romagnoli S, Ronco C. NDT 2019 1-3

Thus, current evidence does not provide a reason to choose a **convective modality over a diffusive** modality in the treatment of patients with AKI.



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Review on RRT in the ICU

A
o SOLUTE A (3 kD)

HEMOFILTRATION

CRRT is <u>strongly recommended over IHD</u> for the management of AKI in patients with acute brain injury, in whom acute perturbations in plasma solute concentration may <u>exacerbate intracranial hypertension</u> which, in combination with <u>systemic hypotension</u>, may lead to <u>critical cerebral hypoperfusion</u>.

Davenport A. Hemodial Int 2010; 14(Suppl 1):S27–S31

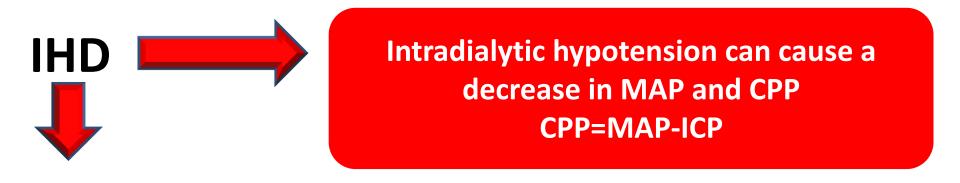


CRRT in Acute Brain Injury

Dialysis disequilibrium syndrome

Conventional IHD may <u>exacerbate</u> the reduction in cerebral perfusion and increase cerebral edema.

Rapid urea removal from the plasma and water shift to the intracellular compartment can worsen brain edema.



Increase ICP by compensatory cerebral vasodilation.

Macedo E et al. Am J Kidney Dis. 2016;68:645-65

CRRT in Acute Brain Injury

IHD should be avoided because it is associated with a more significant increase in ICP compared to CRRT.

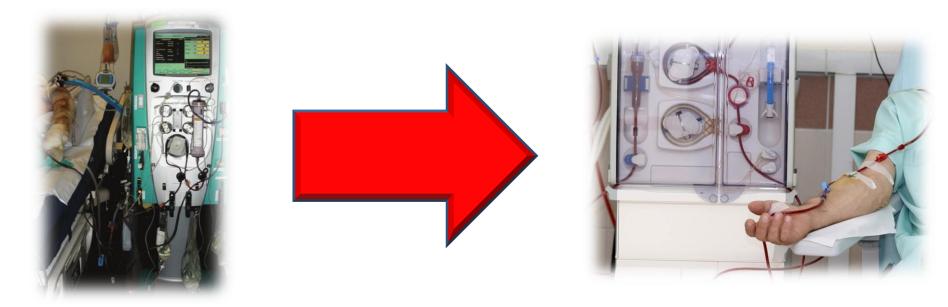


"CRRT, rather than intermittent RRT, for AKI patients with acute brain injury or other causes of increased intracranial pressure or generalized brain edema. (2B)"

Kidney Disease Improving Global Outcomes: Kidney Int Suppl 2012, 2.

Macedo E et al. Am J Kidney Dis. 2016;68:645-65

WHAT IS THE OPTIMAL RRT MODALITY FOR PATIENTS WITH AKI?



- Many patients <u>transition</u> from one modality to another during the course of their AKI, usually from CRRT to IHD as their hemodynamic instability resolves.
- Increasingly, hybrid modalities that provide RRT for 6–18 hours per day, collectively referred to as prolonged intermittent RRT (PIRRT), are used either as an initial modality or as a transition between CRRT and IHD

Edrees F et al. Adv Chronic Kidney Dis. 2016;23(3):195-202. Rachoin JB & Weisberg LS. CCM 2019 Feb 13





CRRT: concetti di base 3

Grazie !

Stefano Romagnoli