

CRRT vs IHD in ICU - a nephrologist point of view

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INTRODUCTION

- Acute kidney injury (AKI) is a part of the multiple organ dysfunction syndrome in critically ill patients
- AKI develops in 30-60% of patients followed in the ICU, approximately 20% of patients with AKI, and approximately 5% of all ICU patients require renal replacement therapy (RRT)
- The treatment of AKI is based on both conservative measures and timely use of RRT

Which type of renal replacement therapy for AKI treatment in ICU?

CRRT?

IHD?

SLED?

Table 1. Characteristics of different modalities of renal replacement therapy for acute kidney injury in ICU

Characteristics	Intermittent hemodialysis	Sustained low-efficiency dialysis	Continuous renal replacement therapy
RRT duration	Intermittent, three times a week, 3–5 h each session	Intermittent, daily, 6–8 h each session	Continuous until filters are clotted or no need for RRT
Doses	Based on URR and Kt/V_{urea} for quantification		~25 ml/kg/h
Anticoagulation	Short exposure	Extended exposure	Often continuous exposure

CRRT

The most appropriate modality in ICU for patients with:

- brain edema

- increased intracranial pressure

- acute brain injury

- fulminant hepatic failure

- head trauma

Wang AJ, Bellomo R. Curr Opin Crit Care 2018; 24: 437-42.

Macedo E, Cerda J. Semin Dial 2021; 34:423-31.

Karkar A, Ronco C. Ann Intensive Care 2020; 10:32.

IHD

- Use of higher flow rates than CRRT to maintain fluid, electrolytes and acid-base balance
- For hemodynamically stable patients
- More suited for patients who require faster removal of uremic toxins and control of electrolyte and acid–base disturbance
- The same IHD machine available for several patients in a same day
- Associated with an increased risk of hypotension, renal ischemia

SLED

- Is a modified form of IHD
- An example of hybrid therapy and extended daily dialysis
- In comparison with IHD:
- Lower dialysate (100-350 ml/min)
- Lower blood flow rate (100-200 ml/min)
- Prolonged session duration than traditional IHD (6-12h)

Table 1 | Treatment parameters for current and previous SLED studies

Author (reference)	Kumar <i>et al.</i> ⁴	Marshall <i>et al.</i> ⁵	Marshall <i>et al.</i> ⁶	This study
Treatment name	EDD	SLED	SLEDD- <i>f</i>	SLED
Hours/day	7.5	12	8	8
Days/week	6-7	6-7	4-7	6
Blood pump speed (ml/min)	200	100	300	200
Dialysate flow (ml/min)	300	200	200	350
Replacement fluid (ml/min)	—	—	100	17

EDD, extended daily dialysis; SLED, sustained low-efficiency dialysis; SLEDD-*f*, sustained low-efficiency daily diafiltration.

Table 3 | Daily and weekly cost of SLED and CRRT

	SLED (\$)	CRRT citrate (\$)	CRRT heparin (\$)
Supply cost/day	69.75	402.80	334.95
HD RN cost/day	168.75 ^a	37.50	37.50
Total cost/day	238.50	440.30	372.45
Total cost/week	1431	3089	2607

CRRT, continuous renal replacement therapy; HD, hemodialysis; RN, registered nurse; SLED, sustained low-efficiency dialysis.

^aNote: Based on one HD nurse treating two patients.



STUDIES



CRRT improves renal recovery from acute renal failure



93 ICU pts with AKI

TABLE I Demographics*

	CRRT (<i>n</i> = 65)	IHD (<i>n</i> = 28)	<i>P</i> value
Time to RRT (hr)	84 (± 80)	68 (± 60)	0.52
Age (yr)	54.7 (± 15.4)	62.6 (± 13.4)	0.02
<i>Gender</i>			
Male	45 (69%)	17 (61%)	0.43
Female	20 (31%)	11 (39%)	
<i>Diagnostic group</i>			
Medical	46 (71%)	17 (61%)	
Surgical	12 (18%)	10 (36%)	0.23
Transplant	7 (11%)	1 (3%)	
APACHE II score	25.1 (± 7.3)	23.5 (± 8.5)	0.37
TISS	47.8 (± 1.3)	37.6 (± 2.0)	0.0001
Mechanical ventilation	65 (100%)	28 (100%)	1.0
Acute lung injury	32 (49%)	6 (21%)	0.01
Admission serum creatinine (µmol·L ⁻¹)	289 (± 217)	410 (± 223)	0.02
Vasoactive drugs required	40 (62%)	10 (36%)	0.02

TABLE III Comparison of indications for RRT mode at initiation of RRT

	CRRT (<i>n</i> = 65)	IHD (<i>n</i> = 28)	<i>P</i> value
Cerebral injury	1 (2%)	0 (0%)	0.51
Hepatic failure	31 (47%)	0 (0%)	0.0001
Dopamine > 5 µg·kg ⁻¹ ·min ⁻¹	18 (27%)	6 (18%)	0.53
Epinephrine	15 (23%)	1 (3%)	0.02
Norepinephrine	29 (44%)	5 (15%)	0.014
Cross over to alternate mode of RRT	18 (67%)	0 (0%)	0.002

CRRT improves renal recovery from acute renal failure

TABLE IV Multivariable logistic analysis of outcomes

	ICU mortality		Hospital mortality		Renal recovery	
	Odds ratio*	P	Odds ratio	P	Odds ratio	P
pH	18 (8, 30)	0.001	8 (5, 10)	0.003		
Age			0.96 (0.93, 0.99)	0.02	0.89 (0.80, 0.99)	0.03
RRT mode					0.04 (0.004, 0.41)	0.006

*Odds ratio (\pm 95% confidence limits). ICU = intensive care unit; RRT = renal replacement therapy.

TABLE V

A) ICU survival vs RRT mode

	Survived	Died	
CRRT	29 (45%)	36 (55%)	<i>P</i> = 0.02
IHD	20 (71%)	8 (29%)	

B) Hospital survival vs RRT mode

	Survived	Died	
CRRT	24 (37%)	41 (63%)	<i>P</i> = 0.24
IHD	14 (50%)	14 (50%)	

C) Renal recovery vs RRT mode

	Recovered	Chronic dialysis	
CRRT	21 (87%)	3 (13%)	<i>P</i> = 0.0003
IHD	5 (36%)	9 (63%)	

Max Bell
SWING
Fredrik Granath
Staffan Schön
Anders Ekblom
Claes-Roland Martling

Continuous renal replacement therapy is associated with less chronic renal failure than intermittent haemodialysis after acute renal failure

Nationwide retrospective cohort study between the years 1995 and 2004. **Follow-up** ranged between 3 months and **10 years**. 2642 pts from 32 Swedish intensive care units included, 2202 pts studied.

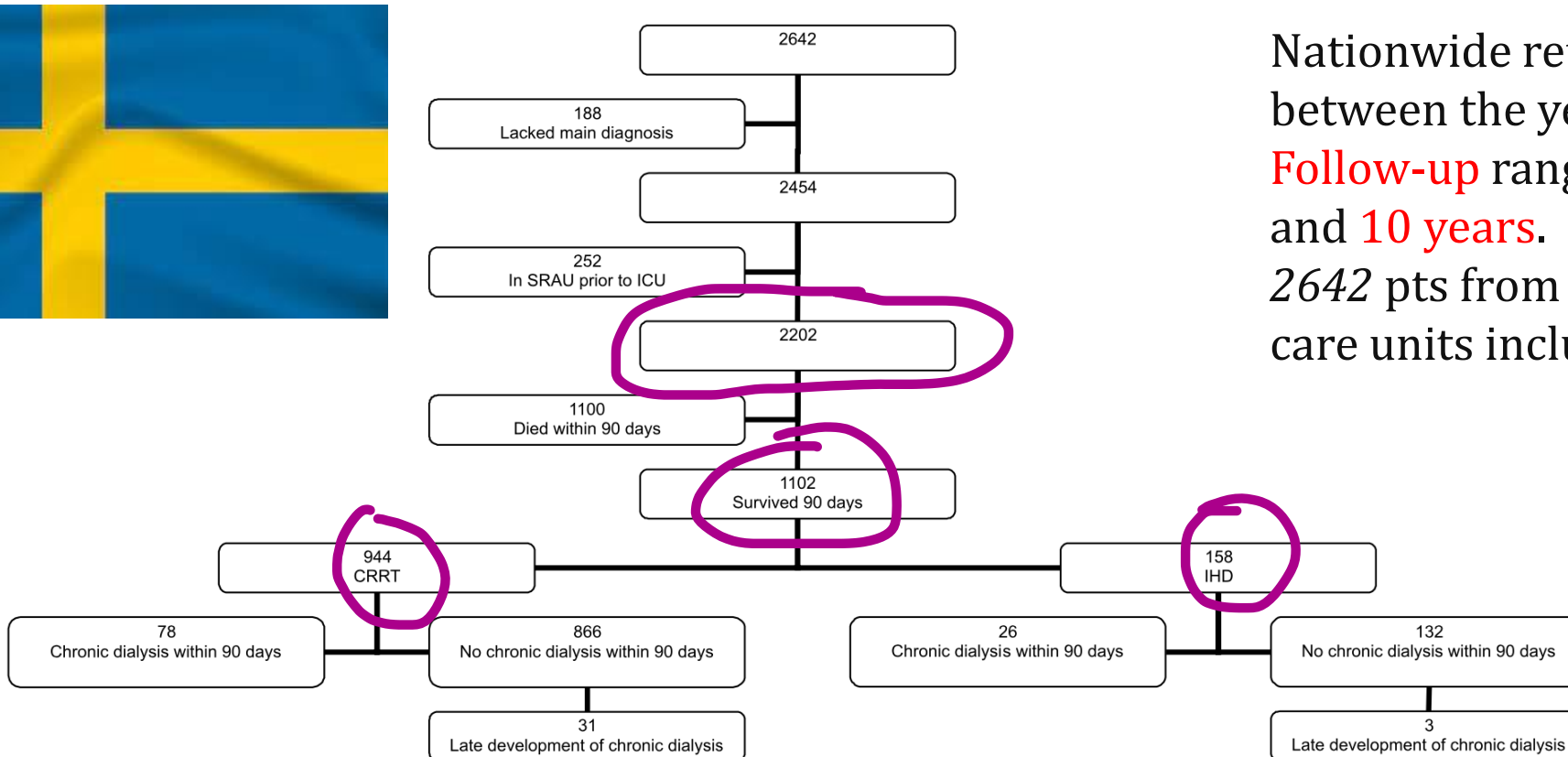


Fig. 1 Flowchart of exclusion criteria and outcome of the cohort

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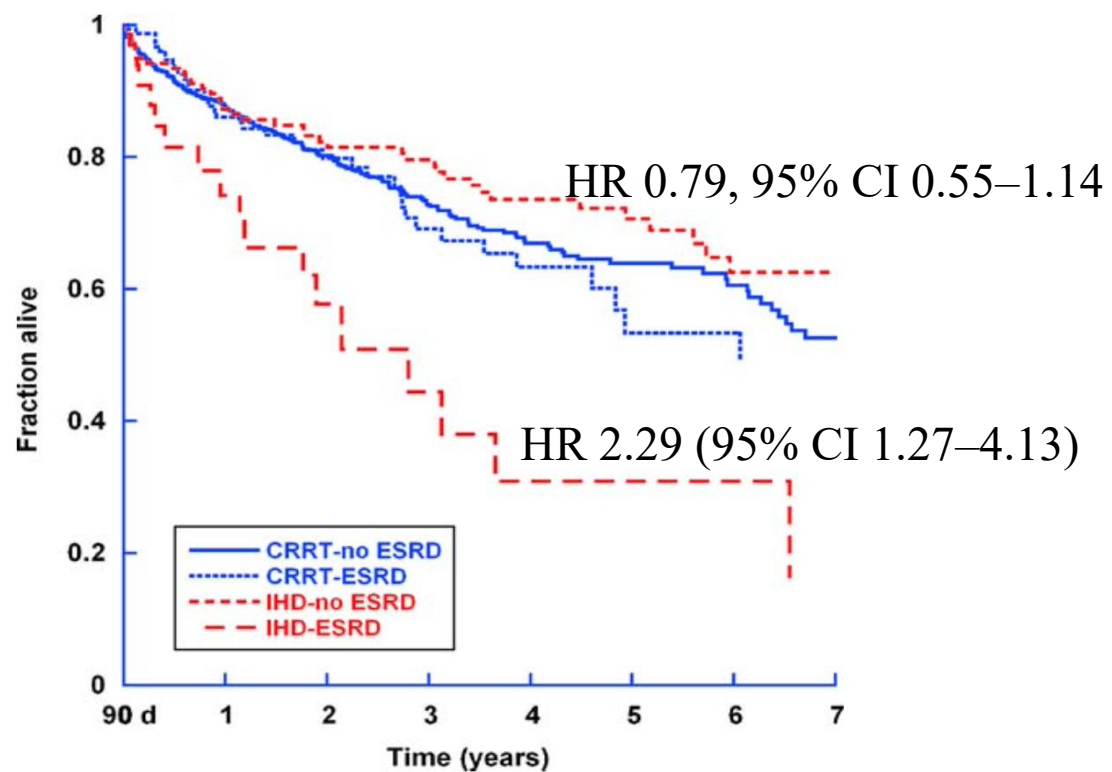


Fig. 2 Kaplan–Meier survival function, all deaths among patients by treatment

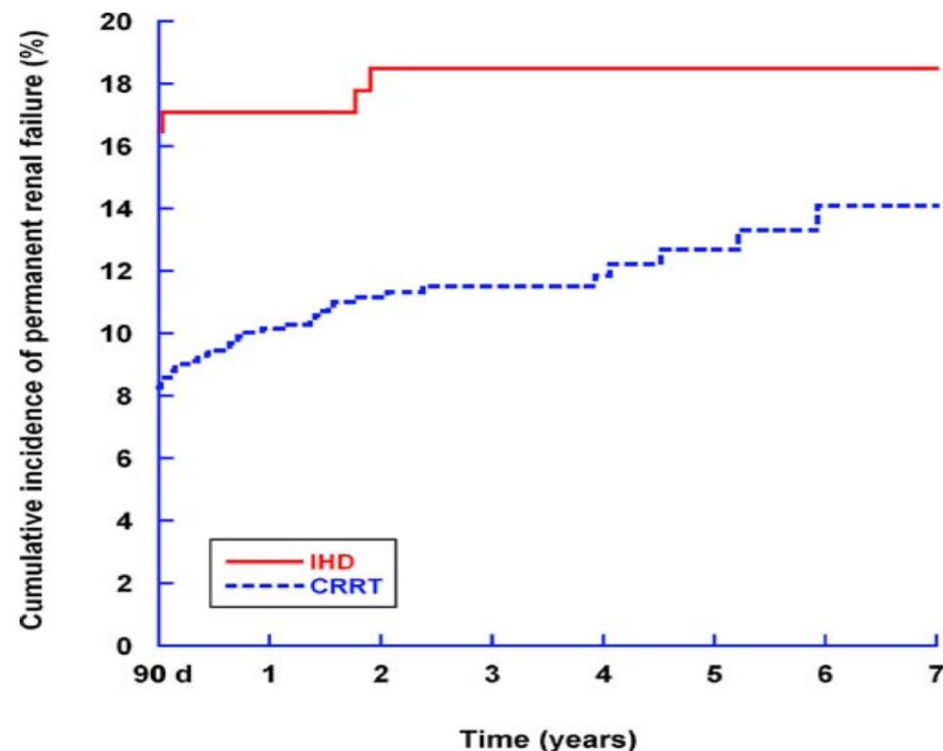


Fig. 3 Cumulative incidence of permanent renal failure among patients surviving 90 days

Patients treated with IHD that ended up with ESRD showed the highest mortality.

A higher risk for ESRD after critical illness for patients treated with IHD than for patients treated with CRRT.

Comparison of the Treatment Efficacy of CRRT and IHD in Patients With AKI admitted to the ICU



120 pts in ICU

Parameters	All Patients (n=120)	IHD (n=80)	CRRT (n=40)	p
Sex: Female; Male	74 (61.7%); 46 (38.3%)	49 (61.3%); 31 (38.7%)	25 (62.5%) 15 (37.5%)	0.894
Age	62.90±13.64	61.41±15.24	65.90±9.13	0.089
SAPS-II	45.05±12.76	43.35±11.11	48.47±15.13	0.038
APACHE II Scores	22.05±6.32	21.06±5.51	24.02±7.38	0.015
SOFA Scores	8.26±2.48	7.91±2.49	8.97±2.35	0.027
Pre-Dialysis				
Mean Arterial Pressure	79.025±15.29	81.75±16.23	73.65±11.62	0.006
Urea (mg/dL)	159.76±66.15	167.18±70.17	144.92±55.13	0.082
Creatinine (mg/dL)	4.09±1.94	4.44±2.09	3.39±1.40	0.005
Sodium (mmol/L)	134.97±6.49	135.08±6.13	134.75±7.24	0.790
Potassium (mmol/L)	4.93±0.94	5.02±1.07	4.73±0.52	0.110
Calcium (mg/dL)	8.36±1.35	8.25±1.42	8.59±1.19	0.198
pH	7.31±0.09	7.29±0.10	7.35±0.07	<0.001
Lactate (mmol/L)	2.33±1.97	2.23±2.22	2.52±1.37	0.448
Bicarbonate (mmol/L)	20.00±4.38	19.59±4.55	20.81±3.95	0.151

Post-Dialysis				
Mean Arterial Pressure	74.49±13.67	75.17±14.67	73.13±11.45	0.442
Urea (mg/dL)	121.08±50.63	115.70±51.21	131.85±48.29	0.100
Creatinine (mg/dL)	3.24±1.46	3.23±1.51	3.27±1.39	0.868
Sodium (mmol/L)	135.80±5.68	136.35±4.73	134.72±7.16	0.140
Potassium (mmol/L)	4.48±0.89	4.48±1.02	4.48±1.02	0.994
Calcium (mg/dL)	8.37±1.01	8.35±1.05	8.43±0.94	0.696
pH	7.32±0.13	7.33±0.14	7.30±0.11	0.338
Lactate (mmol/L)	3.04±3.04	2.88±1.41	3.37±2.10	0.403
Bicarbonate (mmol/L)	21.73±4.14	21.72±4.53	21.75±3.28	0.975
Non-survivor Survivor	80 (66.7%); 40 (33.3%)	53 (66.3%); 27 (33.7%)	27 (67.5%); 13 (32.5%)	0.891
Length of Stay in ICU (days)	12.85±10.23	14.02±10.31	10.50±9.77	0.075

TABLE 1: The demographic, clinical, and laboratory characteristics of the groups during admission to the intensive care unit

Comparison of the Treatment Efficacy of CRRT and IHD in Patients With AKI admitted to the ICU

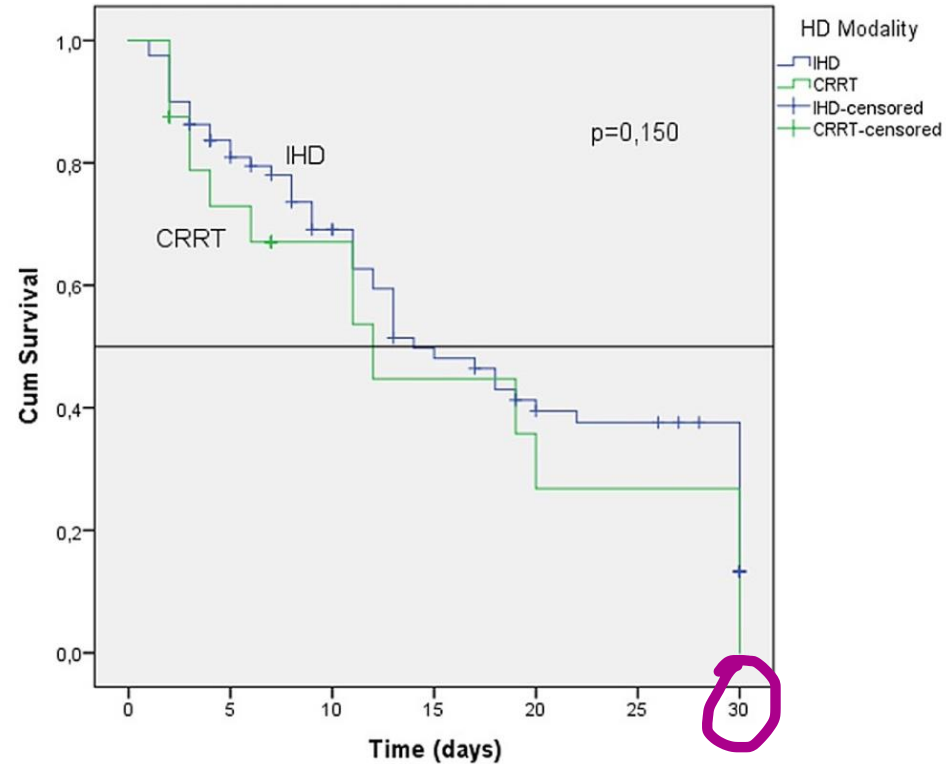


FIGURE 1: Kaplan-Meier survival analysis for hemodialysis modality

Parameters	Univariate			Multivariate		
	OR	95%CI	p	OR	95% CI	p
Age	1.01	0.99 - 1.03	0.183	1.01	0.99 - 1.03	0.116
Sex (Male)	1.19	0.75 - 1.89	0.458	1.32	0.77 - 2.26	0.314
SAPS-II	1.04	1.02 - 1.06	<0.001	1.03	1.01 - 1.06	0.007
APACHE II Scores	1.08	1.04 - 1.11	<0.001	1.05	1.01 - 1.09	0.028
SOFA Scores	1.19	1.10 - 1.30	<0.001	1.13	1.02 - 1.26	0.019
<u>RRT Modality (CRRT)</u>	1.36	0.85 - 2.16	0.198	0.87	0.51 - 1.50	<u>0.616</u>

TABLE 2: Univariate and multivariate Cox Regression analysis results to evaluate 30-day mortality risk

The effect of continuous versus intermittent renal replacement therapy on the outcome of critically ill patients with acute renal failure (CONVINT): a prospective randomized controlled trial

Joerg C Schefold^{1*}, Stephan von Haehling², Rene Pschowski^{1,3}, Thorsten Onno Bender¹, Cathrin Berkmann¹, Sophie Briegel¹, Dietrich Hasper¹ and Achim Jörres¹

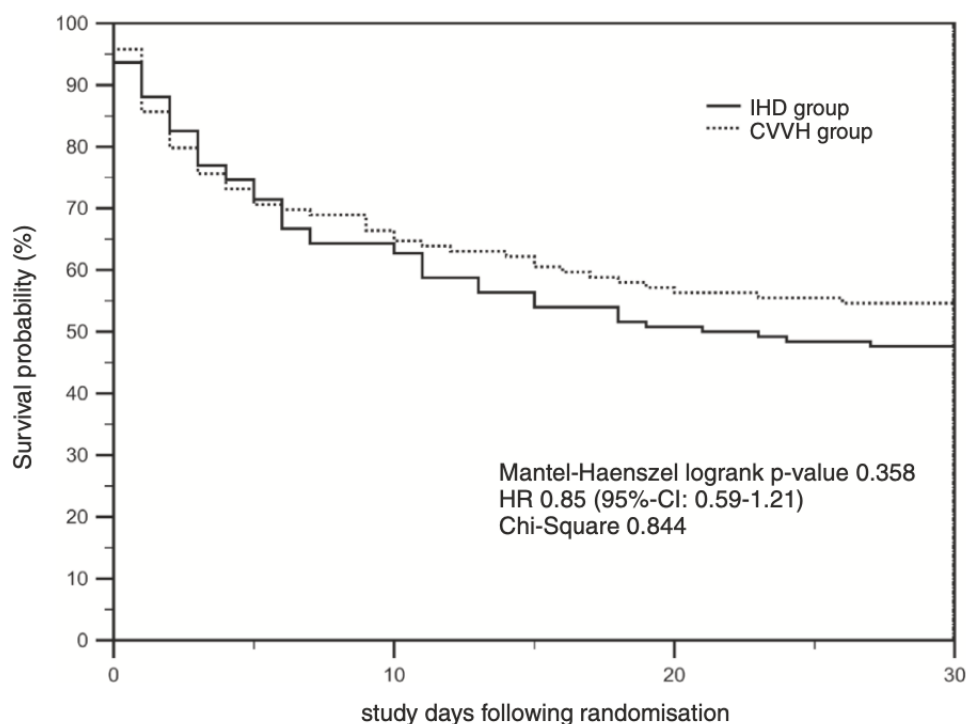

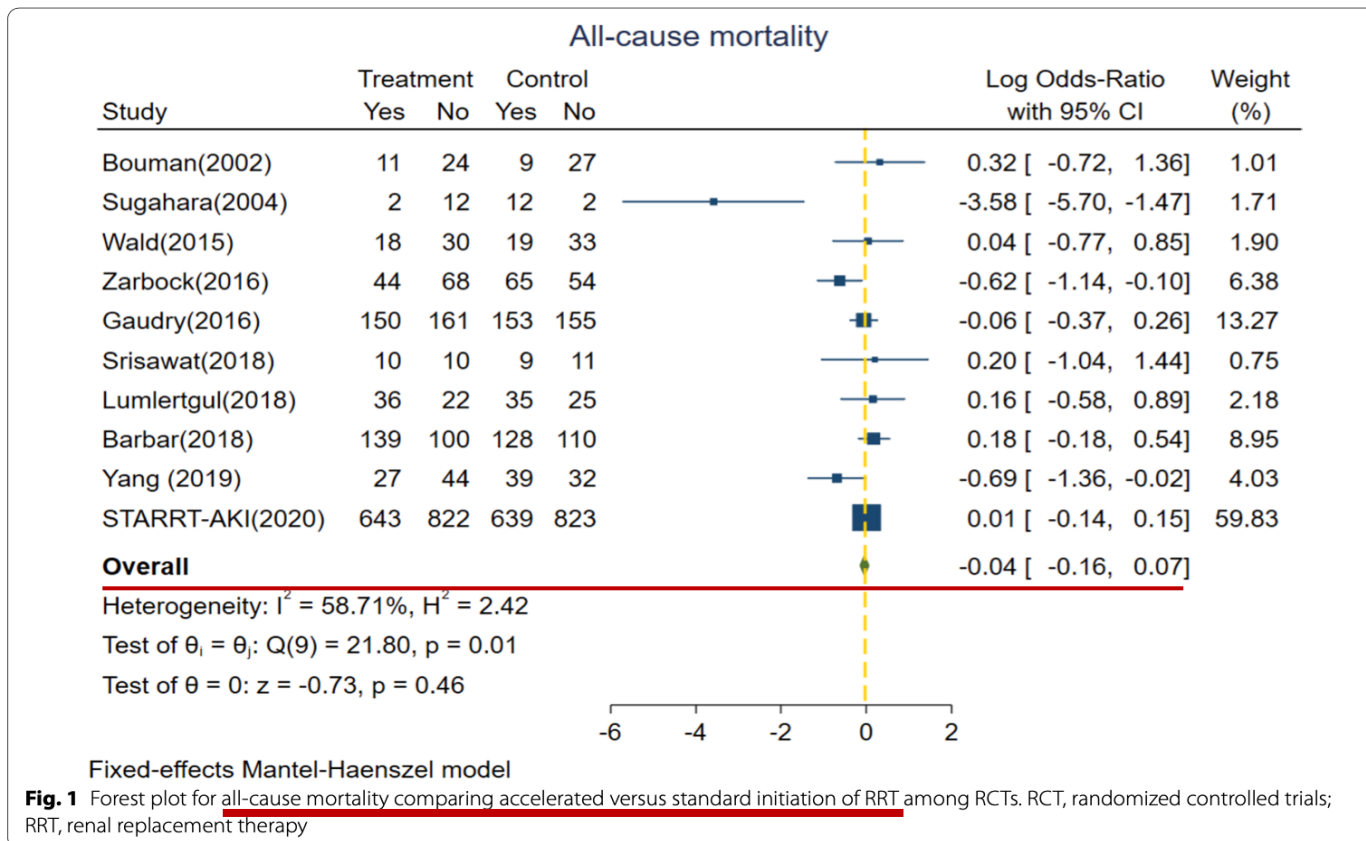


Figure 2 Kaplan-Meier survival estimates for patients randomized to IHD (full line, $n = 128$) and CVVH (dotted line, $n = 122$) are illustrated (total study population). Mantel-Haenszel log-rank P value, hazard ratio (HR) including 95% CI and χ^2 is given.



Accelerated versus standard initiation of renal replacement therapy for critically ill patients with acute kidney injury: a systematic review and meta-analysis of RCT studies

Heng-Chih Pan^{1,2,3,4†}, Ying-Ying Chen^{2,5†}, I-Jung Tsai⁶, Chih-Chung Shiao⁷, Tao-Min Huang⁸, Chieh-Kai Chan⁹, Hung-Wei Liao¹⁰, Tai-Shuan Lai⁸, Yvonne Chueh¹¹, Vin-Cent Wu^{8,12*}  and Yung-Ming Chen^{2,8}



56 studies

10 RCT

4753 critically ill AKI pts in ICU

Critically ill patients with severe AKI would benefit from accelerated RRT initiation only if they were surgical ICU patients or if they underwent CRRT treatment.

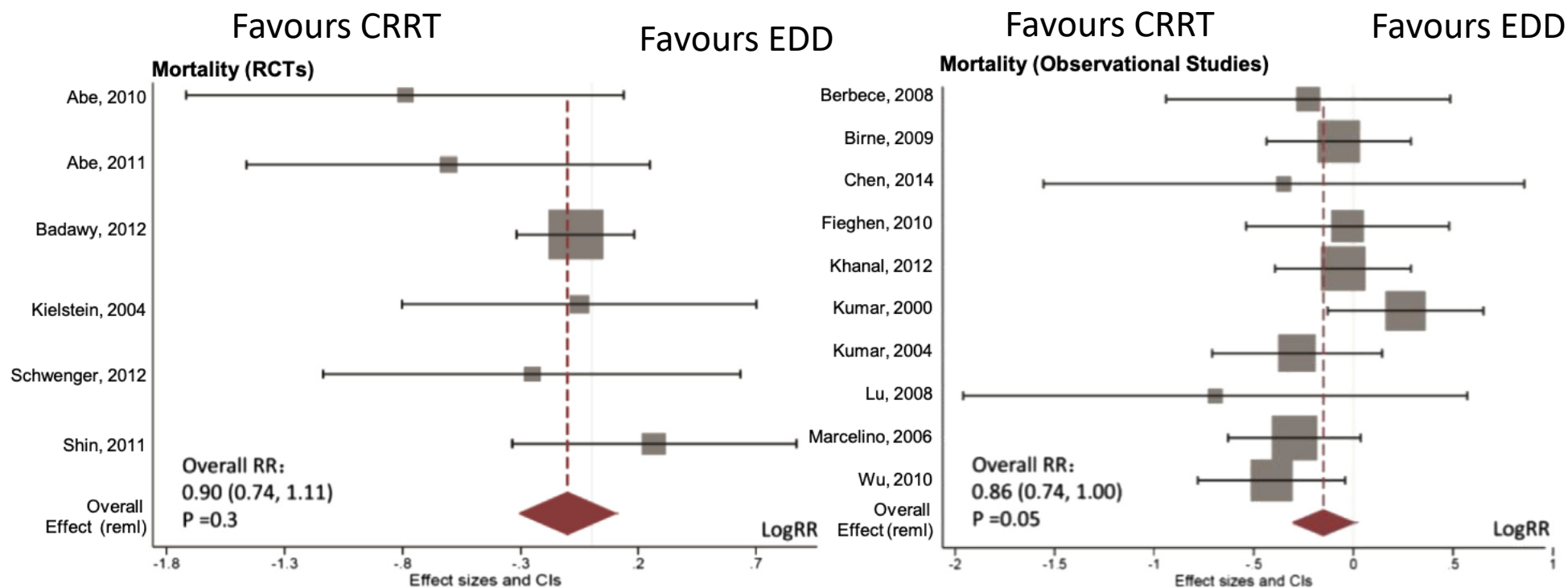
Extended Daily Dialysis Versus Continuous Renal Replacement Therapy for Acute Kidney Injury: A Meta-analysis

*Ling Zhang, MD,^{1,2} Jiqiao Yang, MD,³ Glenn M. Eastwood, MD,² Guijun Zhu, MD,^{2,4}
Aiko Tanaka, MD,² and Rinaldo Bellomo, MD, PhD²*

- primary outcomes were mortality and kidney recovery among patients with AKI
- 7 RCT and 10 observational studies
- from different regions of Asia, North America, Europe, Oceania, and Africa
- The eligible studies were conducted from 2000 to 2014 with a total of **634 patients undergoing EDD** and **574 patients undergoing CRRT**

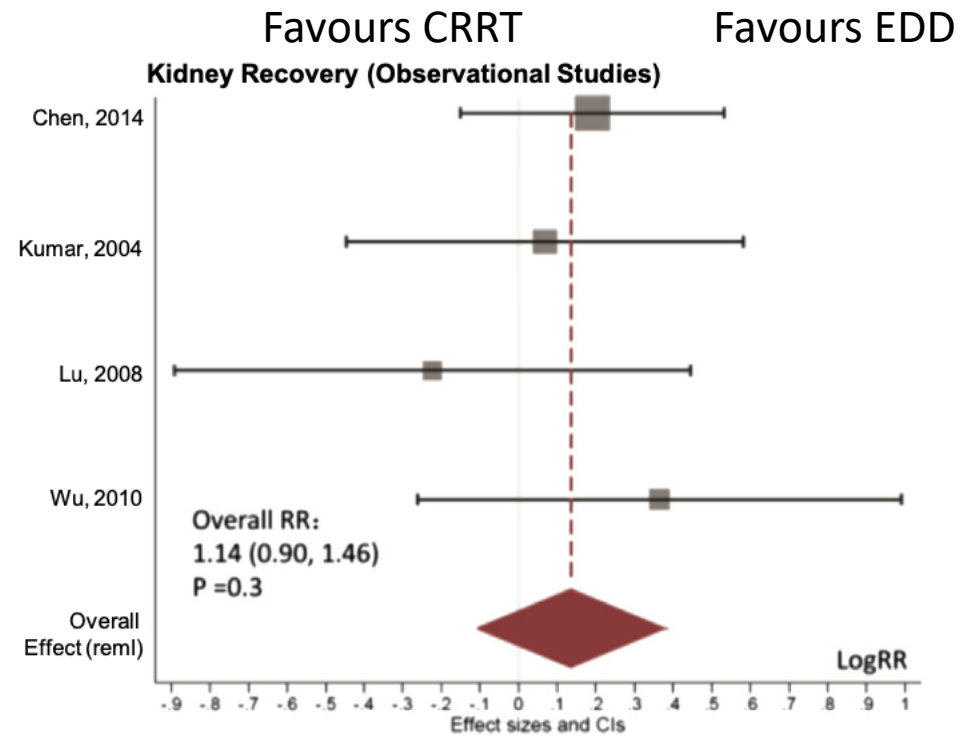
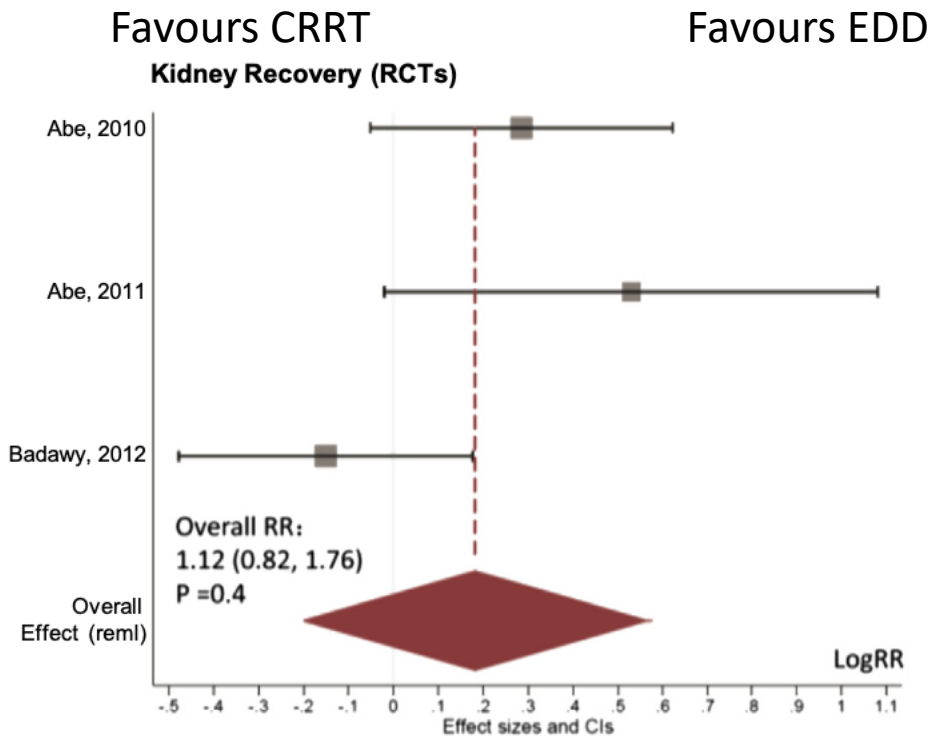
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CRRT vs. IHD as first RRT modality in severe AKI: AKKI and IDEAL-ICU studies

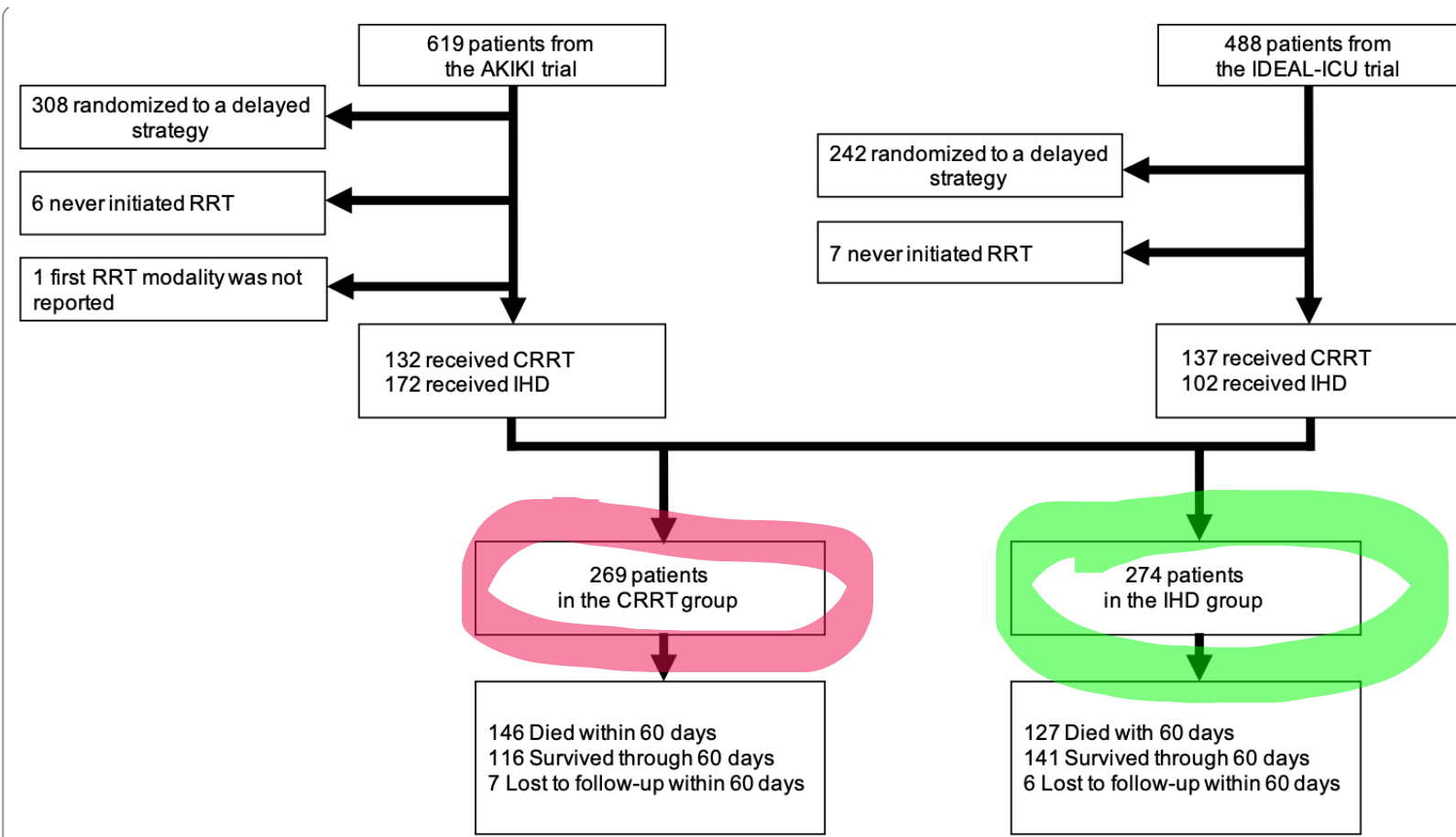
- AKIKI and IDEAL-ICU trials compared an **early RRT initiation** strategy with a **delayed one**
- **Two multicenter RCT** involving **critically ill** patients with severe AKI
- Patients allocated to the **early strategy** received **RRT less than 12 h** after documentation of severe AKI
- No patient was treated with SLED
- Authors analyzed outcomes according to the first modality used

Gaudry *et al.* *Critical Care* (2022) 26:93

Gaudry S, et al. *N Engl J Med.* 2016;375(2):122–33.

Barbar SD, et al. *N Engl J Med.* 2018;379(15):1431–42.

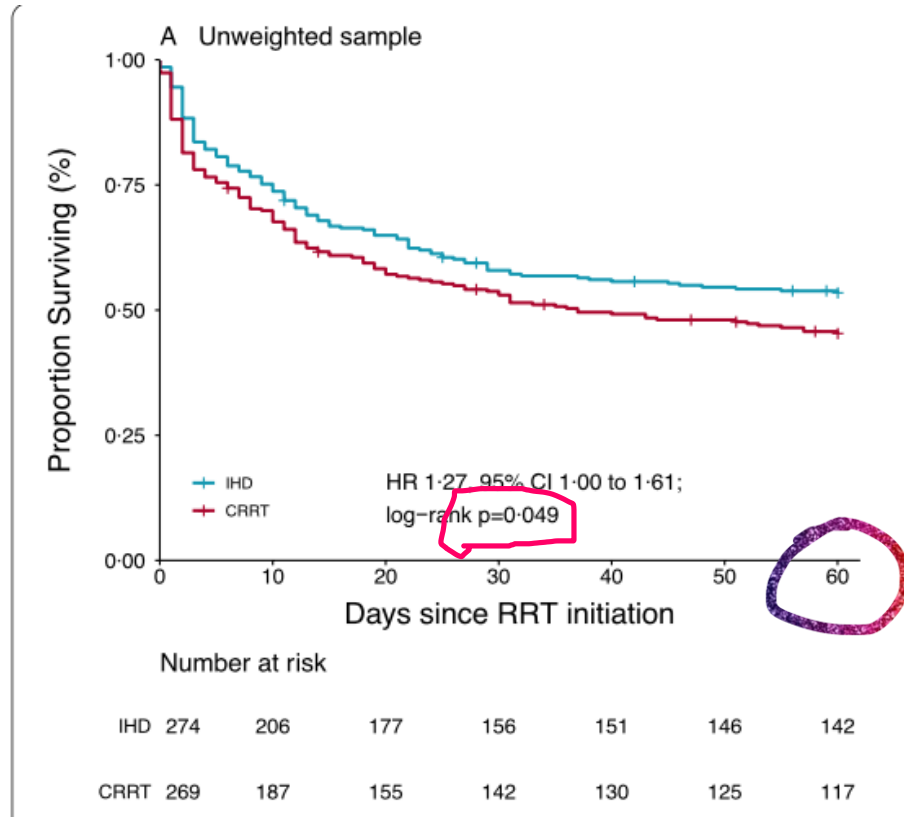
CRRT vs. IHD as first RRT modality in severe AKI: AKKI and IDEAL-ICU studies



Patients have been randomized to receive **either IHD or CRRT within 12 hours** after the documentation of severe AKI.

Fig. 1 Study flowchart

CRRT vs. IHD as first RRT modality in severe AKI: AKKI and IDEAL-ICU studies



- The Kaplan–Meier death rate at day 60:
- 54.7% in the CRRT group
- 46.5% in the IHD group

Primary outcome: probability of survival in the unweighted sample

Gaudry *et al. Critical Care* (2022) 26:93

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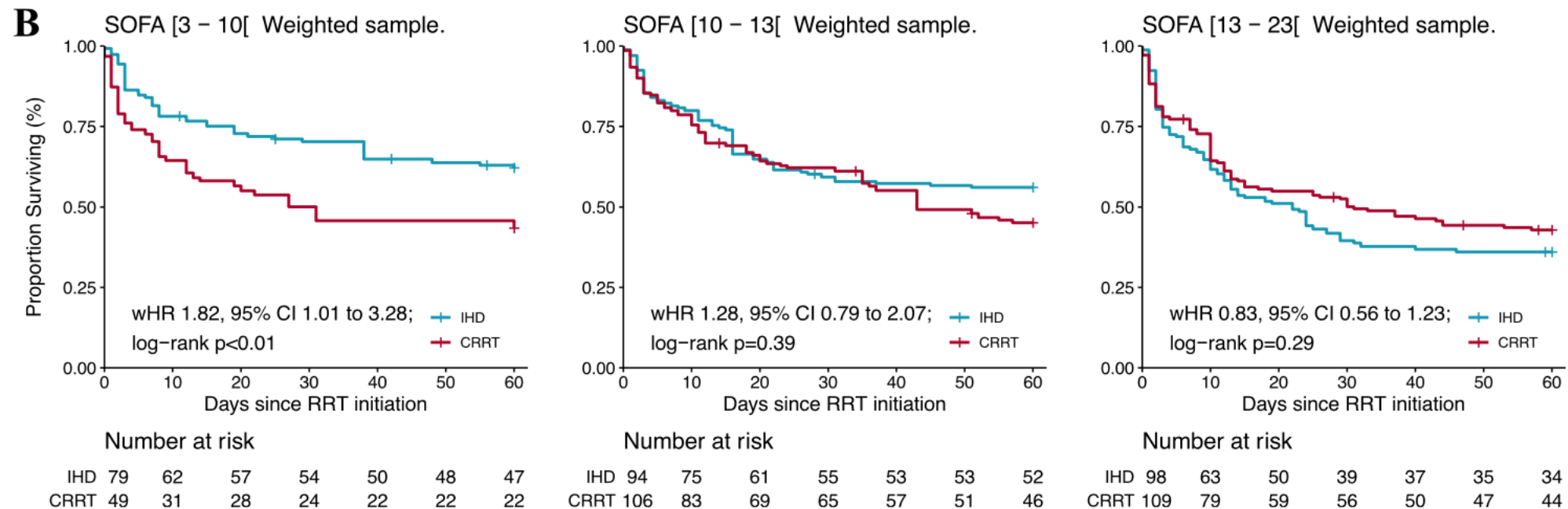


Fig. 3 Treatment effect heterogeneity assessment: Probability of survival in the unweighted samples (A) and in the IPTW samples (B) stratified by thirds of baseline risk (SOFA score). *HR* hazard ratio, *IHD* intermittent hemodialysis, *CRRT* continuous renal replacement therapy

Probability of survival in the weighted samples

Gaudry *et al. Critical Care* (2022) 26:93

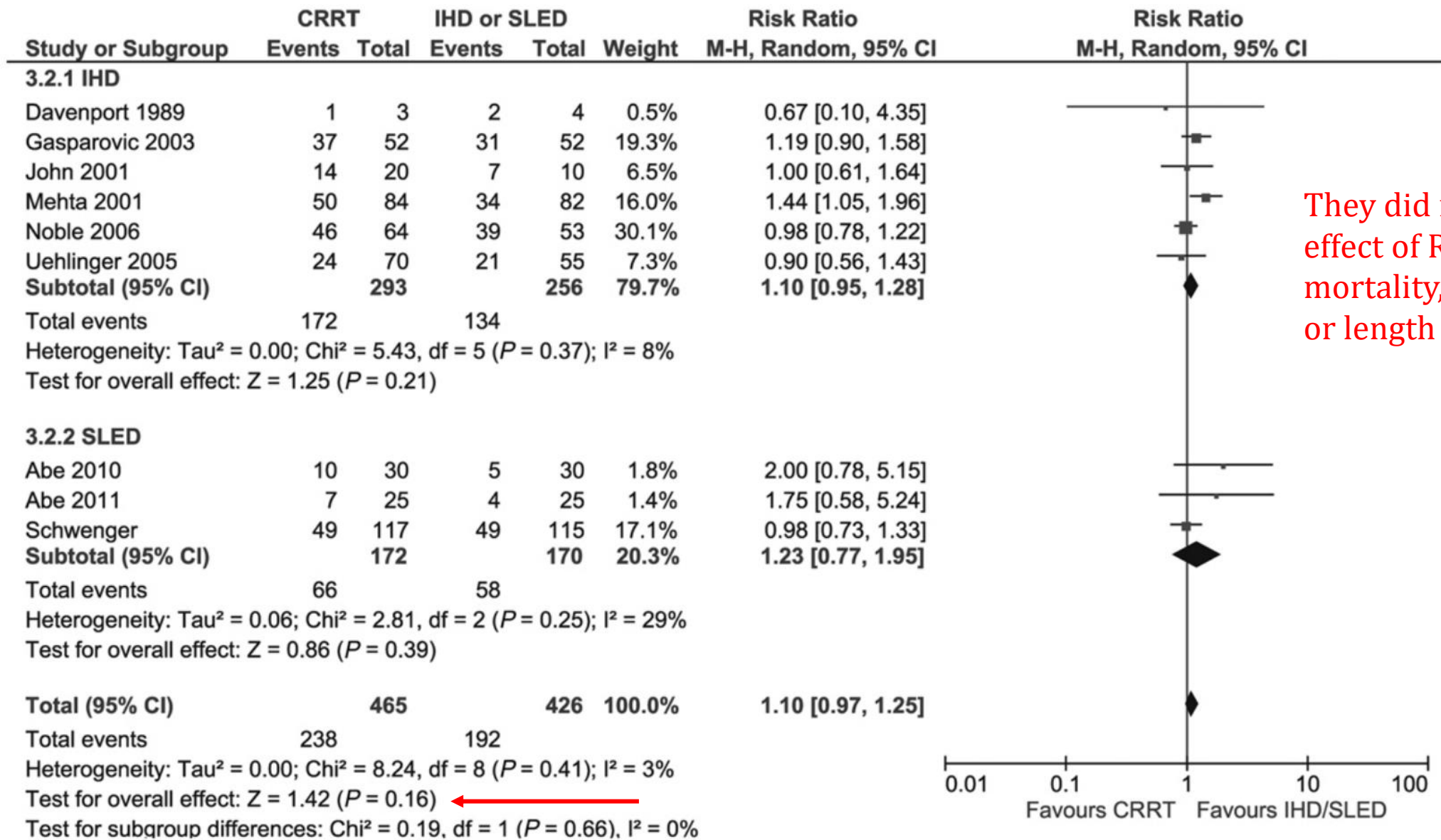
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Barbar SD, et al. *N Engl J Med.* 2018;379(15):1431–42.

Systematic review and meta-analysis of renal replacement therapy modalities for acute kidney injury in the intensive care unit

Danielle M. Nash, MSc ^{a,b,*}, Sebastian Przech, BSc, MDCM ^{b,c,d},
Ron Wald, MDCM, MPH, FRCPC ^{e,f,1}, Daria O'Reilly, MSc, PhD ^{a,g,2}

- 21 eligible studies published from 1989 to 2014 were included
- a total of 5015 participants
- **CRRT vs IHD**: 16 studies, 4539 participants
- **CRRT vs SLED**: five studies (all RCT), 476 participants
- mean age 31 to 69 years;
- over half the studies had a mean participant age over 60 years;
- females proportion ranging from 24 to 43%;



They did not identify any clear effect of RRT modality on mortality, dialysis dependence or length of stay.

Fig. 3. ICU mortality comparing continuous renal replacement therapy and intermittent hemodialysis/sustained low efficiency dialysis.

CHOOSING RRT MODALITY IN ICU- a nephrologist point of view

- Must be patient oriented - patients first!
- Hemodynamic (in)stability, comorbidities?
- Depend of hemodialysis infrastructure
- Depend of dialysis or ICU staff experiences and availability

CONCLUSIONS

- No RRT is ideal for all patients with AKI.
- In the presence of hemodynamic instability in patients with AKI, CRRT is preferable to standard IHD.
- IHD treatment of AKI may delay renal recovery.
- There is lack of solid evidence showing superiority of any mode of RRT in patients with severe AKI in terms of patient survival.



University of Maribor

Faculty of Medicine



THANK YOU FOR YOUR ATTENTION.

