Haemodiafiltration and mortality in end stage kidney disease patients: an individual participant data meta-analysis



The HDF Pooling Project

Sanne Peters, Michiel Bots Bernard Canaud, Andrew Davenport, Muriel Grooteman, Fatih Kircelli, Francesco Locatelli, Francesco Locatelli, Francesco Maduell, Marion Morena, Menso Nubé, Ercan Ok, Ferran Torres, Mark Woodward, Peter J. Blankestijn

RCTs evaluating HDF vs HD

CONTRAST 715 patient, JASN 2012 Turkish trial, 782 patients, NDT 2013 ESHOL, 906 patients, JASN 2013 French, 420 patients, unpublished

Aim of this study:

compare effects of online HDF and standard HD on all cause and cause specific mortality in ESKD patients

Pooling all individual data of three published trials and adding the one unpublished trial

For this analysis additional follow up data on all cause and cause specific mortality was collected.

2793 patients, median follow up 2.5 y

SUMMARY

Online hemodiafiltration in post-dilution mode:

Individual participant data meta-analysis shows:

- 1. Reduction in all cause mortality (Table 1)
- 2. Especially when convection volume > 23 L/session (i.e. 69 L/week) (Table 2)
- 3. No clear side effects
- 4. Mechanism(s): unable to confirm original hypothesis
- 5. Great variability in achieved volume (Figure)

Limitations

Individual trials do not have:

a. sufficient power for sub-group analysis
b. ability to adequately adjust for potential confounders

Methodological flaw: selection bias because of censoring due to non-fatal events in two of the trials (no intention to treat analysis). Complete data in other two trials

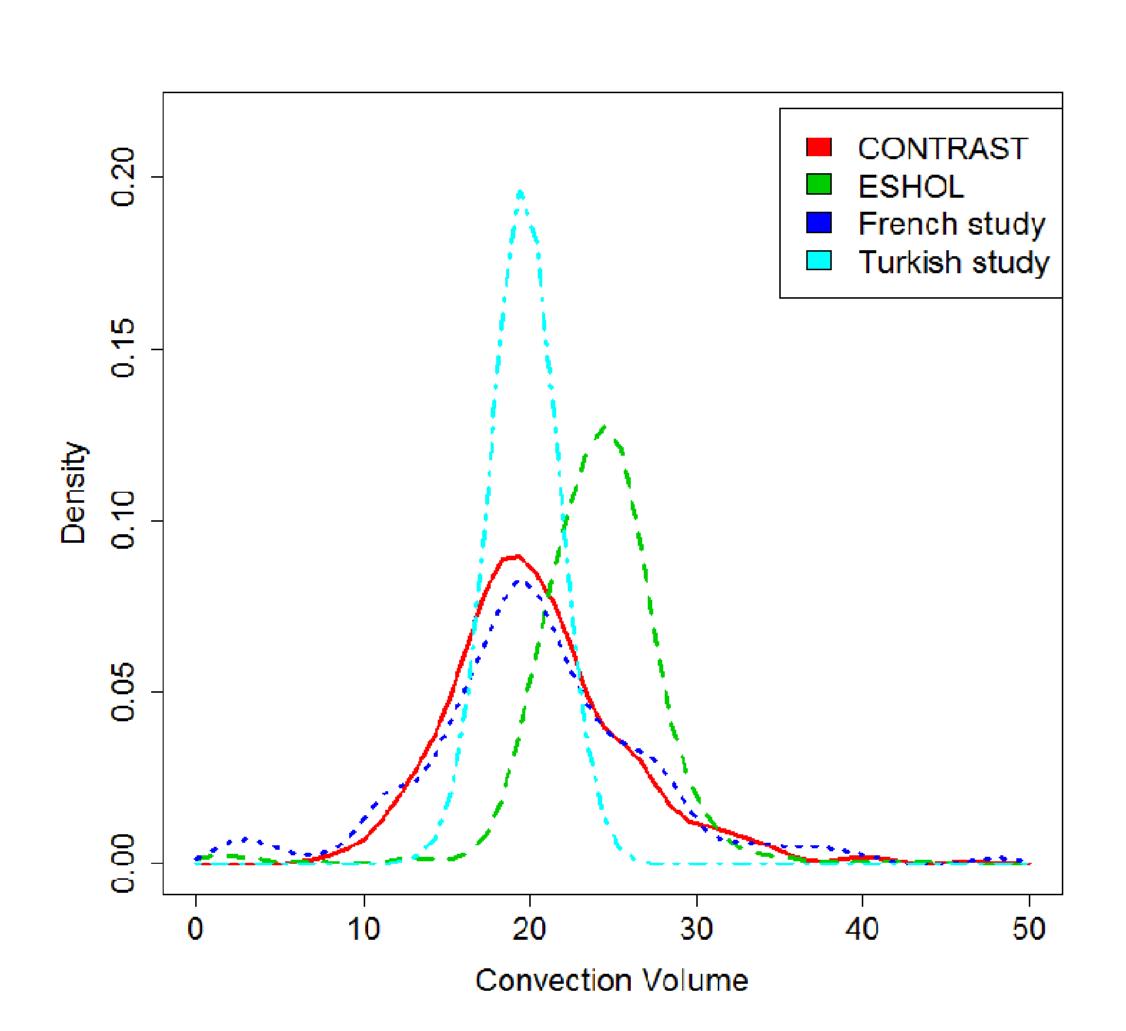


Figure. Distribution of achieved convection volume by study

CONCLUSION

Online hemodiafiltration in post-dilution mode:

- Suggestion of superiority in the same logistical and organizational infrastructure in combination with an apparent lack of side effects, supports the idea of wide spread acceptance
- Time for paradigm shift: dose treatment based on convection volume
- European "end of discussion" trial should study the effects of consistently delivered high dose HDF as compared to standard HD

Table 1. Meta-analysis of all individual data of the 4 RCTs

	Hemodialysis		On line HDF		
	Events	Events/100 PY	Events	Events/100 PY	HR (95% CI)
All-cause mortality	410	12.10	359	10.45	0.86 (0.75; 0.99)
CVD mortality	164	4.84	128	3.73	0.77 (0.61; 0.97)
Infections	77	2.27	73	2.13	0.94 (0.68; 1.30)
Sudden death	56	1.65	56	1.63	0.99 (0.68; 1.43)

Table 2. Risk of mortality by achieved convection volumes (tertiles)

H	Hemodialysis			Online Hemodiafiltration Convection Volume			
		<19	19–23	>23			
All-cause mortality							
Crude	1	0.90 (0.71; 1.13)	0.99 (0.81; 1.20)	0.65 (0.52; 0.82)			
Adjusted	1	0.82 (0.64; 1.04)	1.09 (0.89; 1.33)	0.64 (0.51; 0.81)			
CVD mortality							
Crude	1	0.98 (0.68; 1.40)	0.81 (0.59; 1.11)	0.61 (0.43; 0.88)			
Adjusted	1	0.96 (0.66; 1.39)	0.83 (0.59; 1.16)	0.57 (0.38; 0.84)			
Infections							
Crude	1	1.50 (0.91; 2.48)	0.89 (0.52; 1.51)	0.69 (0.38; 1.25)			
Adjusted	1	1.57 (0.92; 2.67)	0.92 (0.52; 1.63)	0.71 (0.39; 1.30)			
Sudden Death							
Crude	1	1.24 (0.78; 1.97)	1.09 (0.72; 1.67)	0.50 (0.28; 0.88)			
Adjusted	1	1.10 (0.67; 1.82)	1.30 (0.83; 2.03)	0.53 (0.30; 0.96)			











