Outcomes of Warfarin use in Hemodialysis

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Background & Aims

- Patients with end-stage renal disease (ESRD) are at significantly increased risk of major bleeding¹
- There is debate about the safety of warfarin in hemodialysis (HD) patients²
- There is a published potential 4-fold increased bleeding risk in HD patients taking warfarin³

Methods

- We performed a retrospective examination of all patients taking warfarin for any reason whilst on HD on our programme
- Combined time on HD and warfarin was identified and examined
- Outcomes, including bleeding events, thrombotic events and calciphylaxis, were noted
- HD patients regularly have their warfarin dosed by nephrologists, often following telephone advice based on blood tests taken in the dialysis unit
- We questioned if dosing efficacy on the dialysis unit, therefore, contributed to elevated bleeding risk

We aimed to investigate whether time in the therapeutic range (TTR) was optimal in HD patients on warfarin

- Rosendaal linear interpolation formula was used to calculate TTR based on target INR⁴
- TTR is widely used as a marker of safety in oral anticoagulants and a target of 66% is used ⁵
- We also expanded the target range INR for the Rosendaal formula to a wider, and arguably more clinically relevant, therapeutic window of 1.5 - 4
- We noted peak INR results for each patient



- 214 HD patients concurrently taking warfarin were identified over a 10 year period
- A total of 514 patient-years were studied





81 major bleeding events were identified (by ISTH criteria) equalling a bleed rate of 15.9 events per 100patient years (versus a published overall bleed rate of 2.5 per 100 patient years)

AF	61 (29)	TTR	Median (range)
Prosthetic heart valve	42 (19)	Rosendaal for target INR - % tests in range	40 (4.2-96.6)
Thrombus	10 (5)	- % days in range	34.6 (0.8-99.7)
DVT	55 (26)	Beconded for expended IND	
PE	26 (12)	 Rosendaal for expanded INR - % tests in range 	70 (15-100)
Dialysis access patency	8 (4)	- % days in range	68.7 (2.2-100)
Other	12 (5)		

- 8 thrombotic/embolic events were identified
- 6 case of calciphylaxis (calcific uraemic arteriolopathy) were identified, 4 fatal
- Median peak INR was 5.9 overall; 7.4 in patients with a bleed and 3.5 in patients with a thrombotic event
- 75 patients had an INR >5, and 24 >10

Conclusions

- Prevalent haemodialysis patients taking warfarin had a significant bleeding rate
- Thrombotic complications in these patients were low compared with bleeding events
- Of those who had a thrombotic event, the majority had AF and half of them had an embolic event
- Overall, TTR was suboptimal when analyzing for target INR
- Even allowing for expanded acceptable INR targets, TTR was still only just achieved

There is little evidence to support the use of warfarin in hemodialysis patients, but when it is absolutely indicated, this study suggests that TTR in the dialysis population is suboptimal and that supratherapeutic INRs are associated with bleeding risk

References

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- Peak INR and number of INRs above 5 was higher in patients who had a bleed
- Interestingly patients who had a bleed tended to have a higher TTR, although this was not statistically significant
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L7) Dialysis. Epidemiology, outcome research, health services research.

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