INTRAVENOUS LEVETIRACETAM SUPPLETION DURING HAEMODIALYSIS PRESERVED STABLE THERAPEUTIC SERUM CONCENTRATIONS: A CASE REPORT

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Background and Importance

- Levetiracetam is a widely used antiepileptic drug.
- Due to its pharmacokinetic properties, see table 1, it is highly dialyzed during haemodialysis (HD).
- Therefore, it is difficult to preserve stable plasma levels during dialysis and patients starting with HD are often switched to other antiepileptic drugs.
- Information about levetiracetam concentrations in this group of patients are rarely described and show conflicting data.

Aim and Objectives

- We report a case of a 63-yearold woman who started intermittent HD because of renal failure due to diabetic nephropathy.
- She was treated with levetiracetam 250 mg b.i.d. for therapy-resistant focal epilepsy.
- Levels <10 mg/L resulted in frequent seizures, therefore the target values in this patient were set at 10-25 mg/L.

Dialysis settings

- HD sessions lasted 4 hours, trice weekly.
- Fresenius 5008 (Fx CorDiax 1000) HD machine was used.

Table 1: Pharmacokinetic properties of levetiracetam.

Levetiracetam		
Volume of distribution (Vd)	0.5-0.7 L/kg	0
Protein binding	< 10%	H ₃ C NH ₂
Moleculair weight (g/mol)	170.2	N O
Clearance	~100% renal 0.3% via feces	

Materials and Methods

- Additional intravenous doses of levetiracetam were administered during bypass pre-HD, after 2 hours HD and post-HD (see Table 2).
- Levetiracetam concentrations were measured 30 minutes after levetiracetam supplementation.
- Pre-HD samples were measured before the first supplementation dose was given.

Results

Table 2. Levetiracetam supplement and serum concentrations before, during and after haemodialysis sessions.

HD session	1	2	3	4	5	6	7		
Supplemental intravenous dose (mg) of levetiracetam									
Pre-HD				<u>250</u>	250	250	<u>500</u>		
After 2 hours HD	250	250	250	250	250	250	250		
Post-HD	250	250	250	250	250	250	250		
Levetiracetam plasma concentrations (mg/L)									
Pre-HD	18	20	19	18	20	10	19		
After 2,5 hours HD	8	10	10	13	12	17	15		
Post-HD	8	8	17	11	10	9	13		

- Plasma concentrations remained most stable with suppletion doses of 500-250-250mg.
- No seizures or absences have occurred during dialysis and also not on non-dialysis days since the doses were set at 500-250-250mg.

Conclusion and Relevance

- HD showed to eliminate levetiracetam significantly.
- In this case, intravenous levetiracetam suppletion during HD safely preserved stable levetiracetam plasma concentrations preventing seizures.
- Close monitoring of plasma concentrations is recommended to determine the appropriate supplemental dose to maintain therapeutic levels.
- Treatment with anticonvulsant drugs that are not highly eliminated by HD are preferred, e.g. valproate.

Suggestions

 Continuous levetiracetam supplementation during HD could be of further interest.

References

 RenalDrugDatabase, Levetiracetam, accessed 21-06-2023, via: https://renaldrugdatabase.com/monographs/levetiracetam

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