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Predicted environmental concentrations of carbamazepine, oxcarbazepine and their main metabolites in a coastal system

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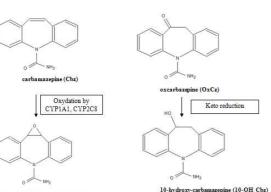
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Hydroxylation





ne-10.11-epoxyde (Cbz Ep

Hydrolysis by Glucurono-conjugation

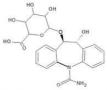


Fig.1: Main metabolic pathways of carbamazepine and oxcarbazepine

Introduction

Pharmaceuticals are widely released in aquatic environment through treated wastewaters. They reach coastal zone indirectly via streams or directly though marine outfalls however data concerning this contamination in coastal waters are scarce.

Environmental Risk Assessment (ERA) of pharmaceuticals have been conducted mostly in surface waters and has not been performed in coastal zone. The first step of ERA is to evaluate the exposure through predictive environmental concentration (PEC) values.

The aim of this study was to predict the occurrence of some pharmaceuticals in a coastal aera subjected to treated wastewater (TWW) reject through a marine outfall (Fig 2.). Among pharmaceuticals, Carbamazepine, Oxcarbazepine and their main metabolites (Fig 1.) were chosen. CBZ has been proposed as an indicator of wastewater contamination and has been already detected in Mediterranean (Munaron et al., 2011).

Prediction was performed based on local pharmaceuticals consumption recording and a review of pharmacokinetics data. PECs values were estimated in TWW and at the marine outfall and compared with MECs obtained by direct quantification and with POCIS implementation.

Materials and methods

PEC

Medical care consumption data (g of CBZ and OxCBZ sales per month)

Pharmacokinetics data (% of excreted forms: parents compounds and metabolites)

% elimination in STEP, flux effluent

PEC effluent: Sales (g) * % excreted/ % elimination*flux PEC coastal zone: PEC/100 (TGD, 2003)

MEC effluents:

24h homogenate effluents sampling (n=8) Filtration, SPE OASIS HLB, analysis LC-MS (Leclercq et al., 2009)

MEC coastal zone: POCIS

Implementation for one month near the submarine outfall (n=6)

Analysis as described by Munaron et al. (2011).

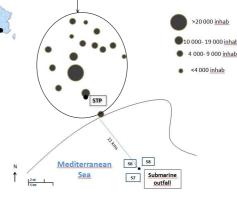


Fig.2: Study area and sampling points (S6-S8)

Results - Discussion

| | 5 | | | -0 |
|-----------------------------|----|--|--|----|
| | 3 | | | -(|
| 9 | 2 | | | |
| Deviation from the mean (%) | 1 | | | |
| ě | -1 | | | |
| E E | -2 | | | |
| ŧ. | -3 | | | |

Fig.3: Deviation from the mean of Cbz and OxCz from January to June 2011

Cbz and Oxcz sale are stable from January to June

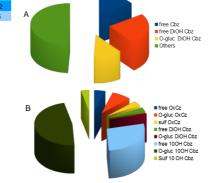


Fig. 4: Metabolits excretion pourcentages of absorbed dose of Cbz(A) and OxCz (B) Cbz and Oxcz are excreted as parent compounds or

metabolites, free or conjugated as glucuronides

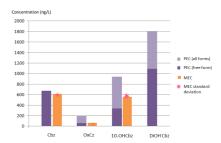


Fig.5: PECs and MECs in treated wastewater effluents All forms (free and conjugated form) considering a total deconjugaison of 0-glucuronides during wastewater treatment

In wastewater effluent, MECs estimation is in good agreement with MECs (Fig5).

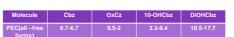


Table 2: PEC in coastal zone (ng/L)



Table 3: concentrations in POCIS in ng/g of sorbent nd not detected

Further studies have to be performed for PEC estimation in coastal area including a hydrodynamic numeric model, which take into account diffusion, advection in seawater.

Conclusion

References

(Fig. 3)

Leclercq, M. et al., 2009. Presence and fate of carbamazepine, oxcarbazepine, and seven of their metabolites at wastewater treatment plants. Archives of environmental contamination and Leclercq, M. et al., 2009. Presence and rate or cardamazepine, oxed basepore, and oxerous members are toxicology 408-415.

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TGD, Technical Guidance Document in support of Commission Directive 93/67/EEC on risk assessment of new notified chemicals and Commission Regulation (EC) No.1488/94 on risk assessment of existing chemicals (1996). Luxembourg: Office for Official Publications of the European Communities; 2003.

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(Fig.4)