Microextraction as sample preparation for new psychoactive substances (NPS) – a greener alternative

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New psychoactive substances continue to appear in the drug market due to the possibility of circumventing legal consequences and generating profit. Consequently the development of new methods is necessary to be able to identify use and quantify the concentrations in biological media. During the last decade an increase in environmental consciousness has led to a wish to reduce the environmental impact of analytical procedures. The term "green chemistry" is used for a set of principles constructed to reduce or eliminate the use or generation of hazardous substances [1]. Sample preparation is a parts of the analytical workflow where the possibility for "greener analysis" is evident [2].

We present three examples of sample preparation for NPS in whole blood using microextraction methods. Analytes were extracted from an aqueous donor phase through a supported liquid membrane (3-4  $\mu$ l organic solvent) into an aqueous acceptor phase compatible with LC-MS/MS. Nitazenes were extracted using 96-well liquid phase membrane extraction, LPME, while tryptamines and synthetic cathinones were extracted using electromembrane extraction, EME. In EME an electric field is the driving force of extraction.

Validation was performed in line with forensic toxicology guidelines, and the methods were evaluated using the AGREEprep tool, developed by Wojnowski et al. [3]. The program gives a score between 0 and 1 for method "greenness". The nitazene method had an intraday precision of  $\leq 10\%$ , interday precision of  $\leq 21\%$ , accuracy  $\leq |8\%|$ , recovery 10–98% and received a greenness score of 0.71. The tryptamine method had intraday precision of  $\leq 16\%$ , interday precision of  $\leq 13\%$ , accuracy  $\leq |6\%|$ , recovery 81-102% and received a greenness score of 0.76. For the synthetic cathinones intraday precision of  $\leq 18\%$ , interday precision of  $\leq 19\%$ , accuracy  $\leq |15\%|$ , and recovery 63-117% were achieved as well as a greenness score of 0.77.

The methods were in line with common requirements for forensic methods, but received very good greenness scores. In comparison a standard method could typically have a greenness score of 0.52 [4].

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