

authentic ephedrine and samples (nos 1 and 6). The spectra of peaks 1 and 2 in the chromatogram of Fig. 2(B and C) matched the spectra of the standard solution of *l*-ephedrine and *d*-pseudoephedrine in the chromatogram of Fig. 2(A).

Conclusion

This simple and convenient HPLC method could determine ephedrine and pseudoephedrine at levels as low as 3 ppm in bulk methamphetamine. As it is very difficult to extract trace ephedrine from the structural analog methamphetamine, this method without any extraction or derivatization procedure may bring benefits for the profiling of high-purity methamphetamine. This developed method was confirmed to be effective for qualitative determination of ephedrine in illicit bulk methamphetamine hydrochloride samples, in which the contents were substantially below the detection limit of the previous method. The configuration of ephedrine is not acquired by this method, but the detection of trace ephedrine may provide valuable information in impurity profiling of high-purity samples. If the content of ephedrine is rich in seized methamphetamine sample, it's favorable to confirm the configuration of ephedrine by our previous method (Makino *et al.*, 2002). Illicit *d*-methamphetamine is prepared only from *l*-ephedrine or *d*-pseudoephedrine, not from *d*-ephedrine or *l*-pseudoephedrine. The ephedrine and pseudoephedrine detected in seized samples may be considered *l*-ephedrine and *d*-pseudoephedrine. Conventional HPLC apparatus is available at most forensic laboratories, so the present method should be widely applicable for identifying trace ephedrine in bulk methamphetamine hydrochloride, and should be helpful for monitoring trends in synthetic methods and precursors used for the illicit production of methamphetamine hydrochloride.

Acknowledgments

I thank the Shiseido Research Center for technical advice on HPLC analysis. This work was done with technical and financial assistance from Professor Tetsuo Nagano at the Graduate School

of Pharmaceutical Sciences, the University of Tokyo, and was also supported by a Health Sciences Research Grant from the Ministry of Health, Labor and Welfare, Japan.

References

- Iwata Y, Inoue H, Kuwayama K, Kanamori T, Tsujikawa K, Miyaguchi H and Kishi T. Forensic application of chiral separation of amphetamine-type stimulants to impurity analysis of seized methamphetamine by capillary electrophoresis. *Forensic Science International* 2006; **161**: 92–96.
- Kanda T, Takezawa K, Kobayashi A, Xin X and Kutsuna H. Capcell Pak C18: ultimately silanol-shielded HPLC column for improved peak shapes of basic compounds under neutral conditions. *LC-GC* 2004; **56**: 43.
- Kurashima N, Makino Y, Urano Y, Sanuki K, Ikehara Y and Nagano T. Use of stable isotope ratios for profiling of industrial ephedrine samples: application of hydrogen isotope ratios in combination with carbon and nitrogen. *Forensic Science International* 2009; **189**: 14–18.
- Luo H, Ma L, Paek C and Carr PW. Application of silica-based hyper-crosslinked sulfonate-modified reversed stationary phases for separating highly hydrophilic basic compounds. *Journal of Chromatography A* 2008; **1202**: 8–18.
- Makino Y, Suzuki A, Ogawa T and Shirota O. Direct determination of methamphetamine enantiomers in urine by liquid chromatography with a strong cation-exchange precolumn and phenyl- β -cyclodextrin-bonded semi-microcolumn. *Journal of Chromatography B* 1999; **729**: 97–101.
- Makino Y, Urano Y and Nagano T. Impurity profiling of ephedrine in methamphetamine by high-performance liquid chromatography. *Journal of Chromatography A* 2002; **947**: 151–154.
- Makino Y, Urano Y and Nagano T. Investigation of the origin of ephedrine and methamphetamine by stable isotope ratio mass spectrometry: a Japanese experience. *Bulletin on Narcotics* 2005; **57**: 63–78.
- Pellati F and Benvenuti S. Determination of ephedrine alkaloids in Ephedra natural products using HPLC on a pentafluorophenylpropyl stationary phase. *Journal of Pharmaceutical and Biomedical Analysis* 2008; **48**: 254–263.
- Remberg B and Stead A H. Drug characterization/impurity profiling, with special focus on methamphetamine: recent work of the United Nations International Drug Control Programme. *Bulletin on Narcotics* 1999; **51**: 97–117.
- Sasaki T and Makino Y. Effective injection in pulsed splitless mode for impurity profiling of methamphetamine crystal by GC or GC/MS. *Forensic Science International* 2006; **160**: 1–10.
- World Drug Report by UNODC, 2010; available from: <http://www.unodc.org/unodc/en/data-and-analysis/WDR-2010.html>

