### 研究论文

高效液相色谱法测定尿中苯的代谢物反,反-粘糠酸 刘黎文宋世震胡霞敏叶方立

武汉科技大学, 湖北 武汉 430080

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摘要 建立了高效液相色谱测定职业苯接触者尿中苯的代谢物反,反-粘糠酸(tt-MA)的方法。该方法采用C18柱进行分离,以冰乙酸-四氢呋喃-甲醇-水(体积比为1:2:10:87)为流动相,以香草酸为内标,于264 nm处进行紫外检测。尿样经2 mo1/L盐酸酸化后用乙酸乙酯进行萃取。结果表明,所建立的标准曲线在tt-MA的质量浓度为0.10~10.00 mg/L时线性关系良好(r=0.9999),加标回收率为95.1%~100.5%,日内和日间测定的相对标准偏差分别为4.0%~9.0%和6.2%~8.8%。应用该法测定职业苯接触者56人和非职业苯接触者24人尿中的tt-MA,结果显示职业苯接触者的尿中tt-MA含量明显高于非职业苯接触者,并与接触的苯的浓度呈线性相关(P<0.01)。该方法灵敏、快速、经济、简便,可用于职业苯接触者的生物监测和毒物动力学研究。

关键词 <u>高效液相色谱法</u> <u>苯代谢物</u> <u>反,反-粘糠酸</u> <u>尿</u> 分类号

# **Determination of Urinary trans, trans-Muconic Acid by High Performance Liquid Chromatography**

LIU Liwen, SONG Shizhen, HU Xiamin, YE Fangli

Wuhan University of Science and Technology, Wuhan 430080, China

#### Abstract

A method for the determination of urinary trans, trans-muconic acid (tt-MA) (benzene metabolite) by high performance liquid chromatography (HPLC) was developed. The separation was carried out on a C18 column (Spherisorb C18, 150 mm×4.6 mm i.d., 5 μm) at 25 °C with glacial acetic acid-tetrahydrofuran-methanol-water (1: 2: 10: 87, v/v) as mobile phase. Urinary sample was acidified by 2 mol/L hydrochloric acid and pretreated by liquid-liquid extraction using diethyl ether. After removal of diethyl ether with a stream of nitrogen, the residue was re-dissolved in 1 mL of mobile phase for HPLC injection. Good linearity was observed within the range from 0.10 mg/L to 10.00 mg/L (r=0.9999) and the detection limit was 0.10 mg/L. The average recoveries for tt-MA were 95.1%-100.5%. Relative standard deviations (RSD) for intra-day and inter-day determinations were 4.0%-9.0% and 6.2%-8.8%, respectively. The method was applied to 56 benzene-exposed workers and 24 controll workers. Urinary tt-MA in benzen-exposed workers were significantly higher than that in the control group. They can be correlated with benzene exposure concentrations (P<0.01). This analytical method for tt-MA is sensitive, rapid, and convenient. It is suitable for monitoring of occupational exposure to benzene and toxic kinetics studies.

**Key words** high performance liquid chromatography (HPLC) benzene metabolite trans, trans-muconic acid urine

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