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## ACCEPTED MANUSCRIPT

# The emergence of low-cost compact MS chromatographic detectors for chemical analysis

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#### 10 Highlights

11 New trends in low-cost compact MS chromatographic detectors for chemical analysis • 12 Recent developments in LC-MS methods using compact MS detectors • 13 Compact MS instruments at half the cost of conventional spectrometers provides excellent • 14 detection sensitivity 15 Use of Microsaic 3500 MiD, Advion Compact and Acquity QDa MS detectors across academia and ٠ 16 industries Use of LC-mini MS for reaction monitoring, biomolecule / high-throughput microplate / impurities / 17 ٠ degradation products / trace / PMIs analyzes 18 19

#### 20 ABSTRACT:

An overview of recent progress in the development of compact mass spectrometers for 21 use as chromatographic detectors in chemical analysis is presented. As the applications of 22 HPLC-MS technologies have grown in recent years there has been a continued expansion 23 of the approach to new user groups. Within the pharmaceutical industry, the recent 24 25 development of small, inexpensive and quiet MS detectors for HPLC has enabled the rollout of this important technology well beyond the initial user base of researchers in 26 drug metabolism and bioanalysis to the direct support of research areas such as discovery 27 chemistry, process chemistry, chemical engineering, manufacturing and formulation 28 sciences, with comparable broadening of the MS user base occurring in other industries 29 and in academia. In this review we survey recent developments and applications ranging 30 from reaction monitoring, biomolecule analysis and high-throughput microplate analysis 31 32 to the identification and analysis of impurities, degradation products and potential mutagens, offering thoughts on current limitations and future directions. 33

*Keywords*: Compact MS; Liquid Chromatography-Mass Spectrometry; Method
 development; High-throughput analysis; Pharmaceutical analysis; Process chemistry
 support.

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