Kenneth J. Fountain and Carsten Wess

## INTRODUCTION

Phenylalanine methyl and other amino acid esters are used as starting material for the production of some active pharmaceutical ingredients (APIs). To ensure the correct and pure isomer is used for synthesis, it is important to perform chiral separations to resolve the enantiomers.

## **CONCLUSIONS**

A UPC $^{2}\text{TM}$  method was developed for the chiral separation of D- and L-phenylalanine methyl esters. The method provides better resolution and 5X the throughput of normal phase HPLC, allowing for high throughput analysis. Due to the low baseline noise observed in UV, the method is capable of detecting down to 500 ng/mL of each enantiomer, which is 0.01% of a 5 mg/mL stock solution. UPC $^2$  provides a rapid method for determining the purity of chiral compounds prior to and during API synthesis.

## INSTRUMENTATION & CONSUMABLES

System: ACQUITY UPC<sup>2™</sup> with photodiode array

(PDA) detection

Column: CHIRALPAK ID, 4.6 x 100 mm, 3 µm

Column temp.: 40 °C

Mobile phase A: CO<sub>2</sub>

Mobile phase B: MeOH with 0.1% NH<sub>4</sub>OH

Isocratic conditions: 90% A, 10% B

Flow rate: 1.5 mL/min

Back pressure: 2500 psi

Detection: UV 210 nm, compensated mode

Injection volume: 4 µL

Sample: 5 mg/mL and 500 ng/mL in isopropanol

with 0.1% triethanolamine

Vials: Waters® Maximum Recovery

Data management: Empower® 3 Software

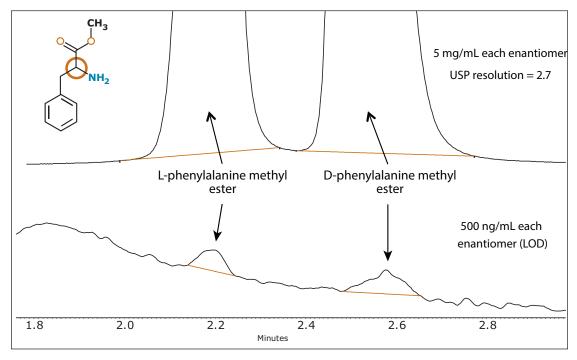


Figure 1. Separation of phenylalanine methyl ester enantiomers using  $UPC^2$ .



## THE SCIENCE OF WHAT'S POSSIBLE.™

RANGE SOUND TO SEE SE SEE SOUND TO SEE SOUND





Waters and Empower are registered trademarks of Waters Corporation. UPC<sup>2</sup>, ACQUITY UPC<sup>2</sup>, and The Science of What's Possible are trademarks of Waters Corporation. All other trademarks are the property of their respective owners.

©2013 Waters Corporation. Produced in the U.S.A. January 2013 720004590EN AG-PDF

Waters Corporation 34 Maple Street Milford, MA 01757 U.S.A. T: 1 508 478 2000 F: 1 508 872 1990 www.waters.com