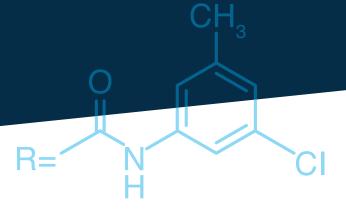
The best separations demand the best columns

INTRODUCING DAICEL'S NEW CHIRALPAK® IK



Companion column to our versatile CHIRALPAK IG

Complementary selectivity with the benefits of our immobilized phases

5 micron available – other particle sizes to follow

| • • • |
|---------------------|
| DAICEL |
| Chiral Technologies |

| Item Code | Description |
|-----------|--------------------------------------|
| 91311 | CHIRALPAK IK, Guard Cartridge (3) |
| 91324 | CHIRALPAK IK, 4.6x150mm, 5µm |
| 91325 | CHIRALPAK IK, 4.6x250mm, 5µm |
| 91335 | CHIRALPAK IK, 10x250mm, 5µm |
| 91337 | CHIRALPAK IK, 10x20mm, 5µm Guard |
| 91342 | CHIRALPAK IK, 20x50mm, 5µm Guard |
| 91345 | CHIRALPAK IK, 20x250mm, 5μm |
| 91355 | CHIRALPAK IK, 50x250mm, 5µm |
| 91372 | CHIRALPAK IK, 30x50mm, 5µm Guard |
| 91375 | CHIRALPAK IK, 30x250mm, 5µm |
| 91394 | CHIRALPAK IK, 2.1x150mm, 5µm |
| 91422 | CHIRALPAK IK SFC, 4.6x50mm, 5µm |
| 91423 | CHIRALPAK IK SFC, 4.6x100mm, 5µm |
| 91432 | CHIRALPAK IK/SFC, 10x50mm, 5µm Guard |
| 91435 | CHIRALPAK IK SFC, 10x250mm, 5µm |
| 91442 | CHIRALPAK IK/SFC, 21x50mm, 5µm Guard |
| 91445 | CHIRALPAK IK SFC, 21x250mm, 5µm |
| 91455 | CHIRALPAK IK/SFC, 50x250mm, 5µm |
| 91472 | CHIRALPAK IK/SFC, 30x50mm, 5µm Guard |
| 91475 | CHIRALPAK IK/SFC, 30x250mm, 5µm |

The Chiral Resolution of β -blocker Acebutolol on CHIRALPAK[®] IK

INTRODUCTION

Acebutolol (Figure 1) is a commonly prescribed β -blocker for the treatment of hypertension and arrhythmias. This is the first reported chiral separation of acebutolol on Daicel polysaccharide chiral stationary phases, and comes on the newest immobilized chiral selector, CHIRALPAK® IK. IK is a cellulose tris(3-chloro-5-methylphenyl)carbamate immobilized on nominal 5 μ m spherical silica gel. The simplicity of the mobile phase (alkane/alcohol) and the robustness of the immobilized phase, are marked improvement over what is current available.

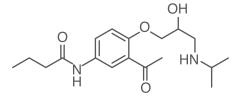


Figure 1: Acebutolol

| Chromatographic Conditions for the Separation of Acebutolol | | |
|---|--|--|
| Column | CHIRALPAK® IK (150mm x 4.6 mm i.d., 5 μ m) Part #: 91324 | |
| Mobile Phase | 70-30-0.1 = Hex-IPA-DEA | |
| Flow Rate | 1.0 ml/min | |
| Detection | UV 254 nm ref. 450 nm | |
| Temperature | 25°C | |
| Sample | 1.2 mg/ml in EtOH | |
| Injection Volume | 5.0 μl | |

EXPERIMENTAL

Acebutolol and Diethylamine (DEA) were purchased from Sigma Aldrich (St. Louis, MO) and used as is. The solvents used were all purchased from Pharmco, were HPLC-grade or higher, and were used as-is. Specifically, the Hexanes (Hex) used contained 95% n-hexane. All screening and optimization were performed on an Agilent 1200 equipped with a quaternary mixing pump, and utilized a DAD.

RESULTS AND CONCLUSIONS

Acebutolol was screening on Daicel's library of immobilized CSPs with starting conditions of 70-30-0.1 = Hex-IPA-DEA. Without any additional optimization, CHIRALPAK® IK provided a baseline resolution of the enantiomers of acebutolol. Figure 2 shows the chromatographic overlay of the screening (with a 20% off-set). The total analysis time was 15 minutes for all methods. The elution times for peak 1 and peak 2 of the separation on IK were 5.65 mins and 7.19 mins respectively.

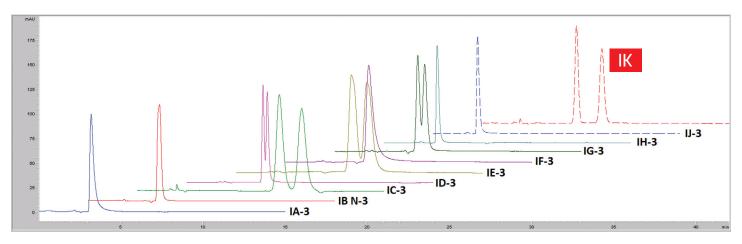


Figure 2: Chromatographic Overlay of the Separation of Acebutolol on Daicel's Library of Immobilized CSPs

Although further optimization could be performed to reduce the analysis time, this was not necessary in this case. It represents the first reported polysaccharide separation of acebutolol, and can be adapted in several ways to other applications, including the use of a longer 250 mm length column, or altering the mobile phase ratio to increase or decrease retention.

