

# PPLICATIONS

## Chiral Separation of Thalidomide with a Lux® i-Amylose-3 Chiral Stationary Phase by HPLC under General Polar Organic Mobile Phase Conditions

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### Overview

In this application, are the details for the separation of a racemic mixture of Thalidomide, using a Lux i-Amylose-3 immobilized polysaccharide chiral LC column and an under polar organic mobile phase conditions. The pharmacological difference between the enantiomers of (S) vs. (R)-thalidomide became apparent as the molecule was marketed for its antiemetic effects, becuase it was later determined that while the (S) form of thalidomide had the desired antiemetic effect, the (R)-(-)-thalidomide enantiomer was causing phocomelia syndrome in infants. This discovery that different enantiomers could have drastically different biological effects/side-effects increased interest in the development of efficient methods to separate the two enantiomers. A general screening polar organic mobile phase, which consisted Acetonitrile with 0.1 Diethylamine. was used on an Agilent® 1100 (Santa Clara, CA) HPLC system, with an injection volume of 10 µL, and a flow rate of 1.0 mL/min. Under these conditions, the Lux i-Amylose-3 was able to separate a racemic mixture of Thalidomide. The Lux i-Amylose-3 chiral selector is a complementary but distinct chiral selectivity, in comparison to i-Amylose-1 and i-Cellulose-5. It combines the 5-position methyl group of the i-Amylose-3 chiral selector, with the 3-position electron-withdrawingchlorinegroupofthe i-Cellulose-5. combined This selectivity with the strong solvent flexibility of the immobilization process promotes the broad enantioselectivity of the Lux i-Amylose-3 column.

### **LC Conditions**

Column: Lux 5 µm i-Amylose-3

Dimensions: 250 x 4.6 mm Part No.: 00G-4779-E0

Recommended Guard: SecurityGuard™ Guard Holder Part No.: KJ0-4282 **Guard Cartridge Part No.:** AJ0-8650

Mobile Phase: Acetonitrile with 0.1 % Diethylamine

Flow Rate: 1.0 mL/min Injection Volume: 10 µL (2 mg/mL) Detector: UV-Vis @ 254 nm Sample: 1. Thalidomide

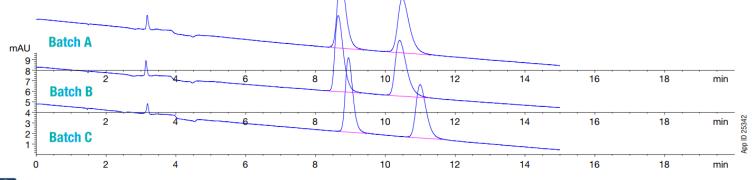
2. Thalidomide

### Lux i-Amylose-3

Amylose tris(3-chloro-5-methylphenylcarbamate)

### **Enantiomers of Thalidomide**

$$\begin{array}{c|c} CH_2 \\ N \\ S \\ CH_2 \end{array} \\ \begin{array}{c} CH_2 \\ N \\ H \end{array} \\ \begin{array}{c} CH_2 \\ R \\ CH_2 \end{array} \\ \begin{array}{c} CH_2 \\ N \\ H \end{array}$$





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