Accepted Manuscript

Rifampin Stability and Solution Concentration Enhancement through Amorphous Solid Dispersion in Cellulose ω -Carboxyalkanoate Matrices

Hale Çiğdem Arca, Laura I. Mosquera-Giraldo, Junia M. Pereira, Nammalwar Sriranganathan, Lynne S. Taylor, Kevin J. Edgar

PII: S0022-3549(17)30428-8

DOI: 10.1016/j.xphs.2017.05.036

Reference: XPHS 839

To appear in: Journal of Pharmaceutical Sciences

Received Date: 14 February 2017

Revised Date: 31 May 2017 Accepted Date: 31 May 2017

Please cite this article as: Arca HÇ, Mosquera-Giraldo LI, Pereira JM, Sriranganathan N, Taylor LS, Edgar KJ, Rifampin Stability and Solution Concentration Enhancement through Amorphous Solid Dispersion in Cellulose ω-Carboxyalkanoate Matrices, *Journal of Pharmaceutical Sciences* (2017), doi: 10.1016/j.xphs.2017.05.036.

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ACCEPTED MANUSCRIPT

- 1 Rifampin Stability and Solution Concentration Enhancement through Amorphous Solid
- 2 Dispersion in Cellulose & Carboxyalkanoate Matrices
- 3 Hale Çiğdem Arca¹, Laura I. Mosquera-Giraldo², Junia M. Pereira¹, Nammalwar Sriranganathan³,
- 4 Lynne S. Taylor², Kevin J. Edgar^{1,4}
- ¹Macromolecules Innovation Institute, Virginia Tech, Blacksburg, VA 24061, USA
- 6 ²Department of Industrial and Physical Pharmacy, Purdue University, IN, 47907, USA
- 7 ³Department of Biomedical Sciences and Pathobiology, Virginia-Maryland College of Veterinary
- 8 Medicine, Virginia Tech, Blacksburg, VA 24061, USA
- 9 ⁴Department of Sustainable Biomaterials, Virginia Tech, Blacksburg, VA 24061, USA

10 Abstract

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Tuberculosis (TB) is a deadly infectious disease; approximately 2 billion people are currently latently infected with the causative agent *Mycobacterium tuberculosis*. Approximately 8 million new active cases and 2 million deaths due to TB are recorded annually¹. Rifampin (Rif) is a vital first line TB treatment drug. Its effectiveness is hampered by the high dose required (600 mg 1x/day) and by its moderate, variable bioavailability. These issues can be explained by Rif instability at gastric pH, limited solubility at neutral pH, polymorphism, and stimulation of its own metabolism. To overcome these obstacles, we developed new cellulose based oral drug delivery systems aiming to increase and make more consistent Rif solubility and bioavailability. Amorphous solid dispersions (ASDs) of Rif with cellulose ω-carboxyalkanoates (cellulose acetate suberate, cellulose acetate propionate adipate, and cellulose acetate butyrate sebacate) were prepared, and compared with crystalline Rif (negative) and carboxymethyl cellulose acetate butyrate ASD (positive) controls. Cellulose ω-carboxyalkanoate

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