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## Biosynthesis of Polyhydroxybutyrate and its Copolymers and Their Use in Controlled Drug

 Release> Abstract: Our aim was to prepare antibiotic loaded rods with biotechnologically produced biodegradable polymers and use them in the treatment of osteomyelitis by providing a high local dose of antibiotic at the infected site. For that purpose, first the production of PHB and its copolymers ( $\mathrm{P}(3 \mathrm{HB}-\mathrm{co}-3 \mathrm{HV}$ ) and $\mathrm{P}(3 \mathrm{HB}-\mathrm{co}-4 \mathrm{HB})$ ) by Alcaligenes latus and Alcaligenes eutrophusin the shake-flask cultures and in a fed-batch fermenter and their purification and characterization were performed. The polymers were then used in the preparation of the sulbactam-cefoperazone loaded rods. To predict the in vivo behavior of the controlled release system, the in vitro release kinetics of the rods were studied in PBS at 37 i C . Release from $50 \%$ w/w loaded $\mathrm{P}(3 \mathrm{HB}-\mathrm{co}-3 \mathrm{HV})$ and $\mathrm{P}(3 \mathrm{HB}-\mathrm{co}-4 \mathrm{HB})$ rods showed that the drug was completely released in less than 3 days. To retard the rate, dip coatings of these rods using the same polymer solution were done and the release profiles were obtained. After coating, cumulative release was about $70 \%$ of its initial content at the end of 12 days. It was concluded that PHB and its copolymers may be a promising alternative to the materials of petrochemical origin in the treatment of osteomyelitis, due to being biodegradable, and eliminating the need for a second operation.

Key Words: polyhydroxyalkanoates, Alcaligenes latus, Alcaligenes eutrophus, osteomyelitis, controlled drug delivery

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