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# Effects of simultaneous internal and external mass transfer and product inhibition on immobilized enzyme-catalyzed reactor

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#### **Abstract**

A mathematical model has been developed for predicting the performance and simulation of a packed bed immobilized enzyme reactor performing lactose hydrolysis, which follows Michaelis—Menten kinetics with competitive product (galactose) inhibition. The performance characteristics of a packed bed immobilized enzyme reactor have been analyzed taking into account the simultaneous effects of internal and external mass transfer limitations. The model design equations are then solved by the method of weighted residuals such as Galerkin's method and orthogonal collocation on finite elements.

The effects of simultaneous internal and external mass transfer coupled with product inhibition have been studied and their effects were shown to reduce internal effectiveness factor. The effects of product inhibition have been investigated at different operating conditions correlated at different regimes using dimensionless  $\beta_{xo}$  (St, Bi,  $\theta$ ,  $\phi$ ). Product inhibition was shown to reduce substrate conversion, and to decrease effectiveness factor when  $\beta_s > \beta_{xo}$ ; however, it increases internal effectiveness factor when  $\beta_s < \beta_{xo}$ . The effectiveness factor is found to be independent of product inhibition at crossover point at which  $\beta_{xo}$  is defined. Effects of St and Bi have been investigated at different kinetic regimes and the results show their effects have a strong dependence on kinetic parameters  $\theta$ ,  $\gamma$  (i.e.  $K_m/K_p$ ) and  $\beta_{xo}$ . The dimensionless residence time at crossover point,  $\beta_{xo}$ , has been correlated with kinetic and mass transfer parameters.

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#### 1. Introduction

Lactose is a disaccharide that occurs naturally in both human and cows milk which accounts for 40% of milk solids. It is widely used in baking and commercial infant-milk formulas. The hydrolysis of lactose, the sugar of milk to glucose and galactose has received much attention in recent years [1,2]. It is used for production of low lactose milk for consumers that suffer from lactose deficiency (70% of the world population is lactose deficient, Carrara and Rubiolo [3]). The hydrolysis product is sweeter and more soluble and biodegradable than lactose and can be used in further biotechnological processes.

The amount of lactose produced annually from whey is about 3.3 million tonnes [3]. It is produced as cheese whey, which is the liquid, separated after milk coagulation. It represents about 90% of the milk volume. The disposal of whey is considered a serious pollution problem facing dairy industry because of its high pollutant content (COD of about 70,000 ppm). Acid hydrolysis of lactose is not favorable because of color formation and fouling of ion exchange resins used in processing. A better alternative is the use of enzymatic method. Enzymatic lactose hydrolysis is carried out by adding β-galactosidase commonly known as lactase to milk, skim milk or whey to hydrolyze lactose prior to pasteurization. Lactase is commercially available and used in large-scale processes. One problem associated with the use of lactase is that complete hydrolysis is difficult to achieve because of product (galactose) inhibition and production of isomer of lactose, allolactose. Several microbial sources of

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#### Nomenclature

- external surface of support per unit volume of
- product concentration in an immobilized  $C_{\mathsf{P}}$ enzyme support particle
- average product concentration in an immobi- $\langle C_{\rm P} \rangle$ lized enzyme support particle
- product concentration in the bulk liquid (reac- $C_{Pb}$ tor phase)
- $C_{Pb0}$ ,  $C_{P0}$  product concentration at reactor inlet
- substrate concentration in an immobilized  $C_{\rm S}$ enzyme support particle
- average substrate concentration in an immobi- $\langle C_{\rm S} \rangle$ lized enzyme support particle
- $C_{\mathrm{Sb}}$ substrate concentration in the bulk liquid
- $C_{\rm Sb0}$ ,  $C_{\rm S0}$  substrate concentration at reactor inlet
- $D_{\mathrm{Sp}}$ ,  $D_{\mathrm{Pp}}$  effective substrate and product diffusivity in an immobilized enzyme support particle
- $D_{Sz}$ ,  $D_{Pz}$  effective substrate and product axial dispersion coefficient
- $K_{\rm e}$ reaction equilibrium constant
- $K_{\rm L}$ mass transfer coefficient
- $K_{L}a$ volumetric mass transfer coefficient
- $K_{LS}$ ,  $K_{LP}$  mass transfer coefficient in substrate and product side, respectively
- intrinsic Michaelis-Menten constant  $K_{\rm m}$
- product inhibition constant  $K_{\mathfrak{p}}$
- L length of the reactor
- radial coordinate of distance in an immobilized r enzyme support particle
- $\tilde{R}_{\rm h}$ dimensionless reaction rate at the surface of the spherical particles
- local product production rate per unit of cat- $R_{\rm P}$ alytic particle volume
- $\langle R_{\rm P} \rangle$ average product production rate
- local substrate consumption rate per unit of  $R_{\rm S}$ catalytic particle volume
- average substrate consumption rate  $\langle R_{\rm S} \rangle$
- time inside reactor
- superfacial fluid phase velocity inside the reac-
- maximum reaction rate per unit of catalytic  $v_{\mathrm{max}}$ particle volume
- reactor radial and axial coordinate x, z

#### Dimensionless variables

- Biot number Bi
- Da Damkohler number
- inverse of the equilibrium constant  $K_{\rm E}$
- P dimensionless product concentration in an immobilized enzyme support particle
- $\langle P \rangle$ dimensionless average product concentration in an immobilized enzyme support particle

$P_{b}$	dimensionless product concentration in the
	bulk liquid
$P_{\mathrm{b0}}$	dimensionless product concentration at the
	reactor inlet
Pe	Peclet number
$ ilde{R}$	dimensionless reaction rate in an immobilized
	enzyme support particle
S	dimensionless substrate concentration in an
	immobilized enzyme support particle
$\langle S \rangle$	dimensionless average substrate concentration
	in an immobilized enzyme support particle
$S_{ m b}$	dimensionless substrate concentration in bulk
	liquid

- $S_{b0}$ dimensionless substrate concentration at reactor inlet
- St Stanton number
- X fractional substrate conversion
- Y reactor yield

#### Greek symbols

- effective diffusivity ratio of substrate and prod- $\alpha_{\rm z}, \alpha_{\rm P}$ uct in axial and interparticle, respectively
- dimensionless residence modulus  $\beta$ ,  $\beta$ <sub>s</sub>
- reactor voidage ε
- Thiele modulus
- dimensionless inhibition modulus γ
- internal effectiveness factor η
- external effectiveness factor  $\eta_{\mathrm{E}}$
- dimensionless Michaelis-Menten constant  $\theta$
- τ dimensionless time
- dimensionless radial coordinate ξ
- dimensionless axial coordinate ζ

β-galctosidase and reactor types have been used for the purpose of economic production of low lactose milk. Lactose hydrolysis in plug flow reactor gives higher conversion compared to continuous stirred tank reactor although the latter has good mixing and lower construction cost.

#### 1.1. Kinetics of lactose hydrolysis

Kinetics of lactose hydrolysis has been studied extensively in the literature [1,2,4]. Michaelis-Menten model with competitive product inhibition by galactose is widely used to describe the hydrolysis [1]. Different types of bioreactor [5,6] and biocatalyst [1,2,7,8] have been investigated for lactose hydrolysis.

#### 1.2. Modeling immobilized enzyme reactor

Enzyme immobilization offers a number of advantages over enzymes in suspension. Immobilization permits the reuse of the enzyme and may provide a better environment

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