

Local Sensitivity Analysis of a Supercritical Extraction Model

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ABSTRACT

This study investigates the process of chamomile oil extraction from chamomile flowers. A parameter-distributed model, consisting of a set of partial differential equations, was used to describe the governing mass transfer phenomena between solid and fluid phases under supercritical conditions using carbon dioxide as the solvent. The concept of quasi-one-dimensional flow was applied to reduce the number of spatial dimensions. The flow of carbon dioxide is assumed to be uniform across any cross-section, although the area available for the fluid phase can vary along the extractor. The physical properties of the solvent were estimated using the Peng-Robinson equation of state. Laboratory experiments were conducted under various, but constant operating conditions of 30–40 °C, 100–200 bar and $3.33 - 6.67 \cdot 10^{-5}$ kg/s. The local sensitivity analysis method was applied to evaluate the robustness of the model parameters by investigating the impact of infinitesimally small changes in the model parameters and controls on the model outputs. This study focuses on analysing the effect of pressure on the model state space and extraction yield. It was found that the model is the most sensitive to the controls if operated close to the critical point.

1. Introduction

Supercritical CO₂ is defined as carbon dioxide that is pressurized and heated above its critical point (31.1 °C, 74 bar). Depending on the operating conditions, the fluid properties such as viscosity and density can vary, which leads to multiple industrial applications of CO₂.

One of the most popular applications of supercritical CO₂ is the extraction of essential oils, as described by many researchers, for example, by Sodeifian and Sajadian [1], Reverchon et al. [2] or Sovova [3]. Traditional methods, such as distillation and organic solvent extraction, are commonly employed but have drawbacks. Distillation involves high temperatures that can lead to the thermal degradation of heat-sensitive compounds. This limitation has increased the popularity of alternative techniques, such as supercritical fluid extraction. Supercritical CO₂ is appealing due to its distinctive properties: it is inflammable, non-toxic and non-corrosive. Supercritical fluids can exhibit both gas- and liquid-like properties, allowing for adjustable dissolving power through changes in operating conditions.

The applications of supercritical carbon dioxide are not limited only to an extraction process but can also be used for impregnation, as described by Weidner [4] or Machado et al. [5]. Impregnation is defined as modifying the properties of bulk substances by physically or chemically binding/adsorbing impregnates to a bulk material or surface, such as the hydrophobization of surfaces. The main advantage of using supercritical CO₂ is that, after depressurization, it desorbs from the surface and evaporates, leaving a solvent-free product. On the other hand, the main disadvantage of using carbon dioxide for impregnation is the low solubility of many drugs of interest. The study of Ameri et al. [6] investigates the loading of lansoprazole into polymers using

supercritical carbon dioxide and examines how various parameters, such as temperature, pressure, and time, affect the drug loading efficiency. The results indicate that increasing any of these parameters enhances drug loading, with temperature having the most significant impact. The study of Fathi et al. [7] explores the use of supercritical carbon dioxide to enhance the bioavailability of ketoconazole by impregnating it into water-soluble polymers, specifically polyvinylpyrrolidone and hydroxypropyl methylcellulose. Utilizing a Box-Behnken design, the researchers optimized the impregnation process by varying pressure (150–250 bar), temperature (308–328 K), and time (60–180 minutes), achieving increase in drug loadings ranging

Another application of supercritical CO₂ is nanoparticle formation, as investigated by Padrela et al. [8], Franco and De Marco [9] or Sodeifian et al. [10]. Supercritical carbon-dioxide-assisted technologies enable the production of different morphologies of different sizes, including nanoparticles and nanocrystals, by modulating the operating conditions. Supercritical fluid-based processes have advantages over techniques conventionally employed to produce nanosized particles or crystals, such as reduced use of toxic solvents. Moreover, the CO₂ is removed from the final product by simple depressurization. Sodeifian and Sajadian [11] investigates the solubility of Letrozole, a poorly water-soluble anticancer drug, in supercritical carbon dioxide with and without menthol as a solid co-solvent. The addition of menthol increased Letrozole's solubility by 7.1 times compared to supercritical CO₂ alone. Using the rapid expansion of supercritical solutions with solid co-solvent method, the average particle size of Letrozole was reduced to the nanoscale. Temperature was found to have the most significant impact on nanoparticle size reduction, while pressure had the least effect. The study of S. Ardestani et al. [12] explores the preparation of phthalocyanine green nano pigment using supercritical carbon dioxide as an antisolvent. The researchers employed the gas antisolvent technique to

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achieve nano-sized particles of the pigment. This method offers a potential approach for producing nano pigments with enhanced properties. The findings contribute to the development of advanced materials in the field of pigments and coatings. The study of Sodeifian and Sajadian [13] investigates the production of amiodarone hydrochloride nanoparticles using an ultrasonic-assisted rapid expansion of supercritical solution into a liquid solvent method. By optimizing parameters such as pressure, temperature, and polymeric stabilizer concentration, the researchers achieved significant particle size reduction. Characterization techniques confirmed the successful formation of nanoparticles with improved properties.

This study investigates the extraction of essential oil from chamomile flowers (*Matricaria chamomilla* L.) via supercritical fluid extraction techniques and the modelling of this process. Chamomile is a medicinal herb widely cultivated in southern and eastern Europe — in countries such as Germany, Hungary, France and Russia. It can also be found outside Europe, for instance in Brazil, as discussed by Singh et al. [14]. This plant is distinguished by its hollow, bright gold cones, housing disc or tubular florets and surrounded by about fifteen white ray or ligulate florets. Chamomile has been used for its medicinal benefits, serving as an anti-inflammatory, antioxidant, mild astringent, and healing remedy. Extracts of chamomile are widely used to calm nerves and mitigate anxiety, hysteria, nightmares, insomnia and other sleep-related conditions, according to Srivastava [15]. Orav et al. [16] reported that oil yields from dried chamomile samples ranged from 0.7 to 6.7 mL/kg. The highest yields of essential oil, between 6.1 and 6.7 mL/kg, were derived from chamomile sourced from Latvia and Ukraine. In comparison, chamomile from Armenia exhibited a lower oil content of 0.7 mL/kg.

The literature offers various mathematical models to describe the extraction of valuable compounds from biomass. Selecting a process model is case-dependent and requires analysis of each model's specific assumptions about mass transfer and thermodynamic equilibrium.

Depending on the needs one of two approaches can be considered while developing a mathematical model. A model, which is based on a multiple regression can be used if the relation between inputs and outputs can be simplified. Sodeifian et al. [17] investigated an influence of pressure, temperature and particle size on the extraction efficiency of oil from *dracocephalum kotschy boiss* seed. The second order polynomial model was applied to obtain the corresponding response surface and to identify the optimum operating conditions. The study of Sodeifian et al. [18] investigates the of essential oil from *Eryngium billardieri*, focusing on optimizing extraction conditions and developing a mathematical model based on the f the second order polynomial to predict the process yield. The researchers employed a simulated annealing algorithm to optimize parameters such as pressure, temperature, and extraction time, aiming to maximize oil efficiency.

Alternatively, a more complex model can be derived and applied to cover not only the input-output relations, but the phenomena occurring in the system. This approach allows for more detailed representation the system behaviour, but requires better understanding of the mass transfer and more rigorous experiments.

Goto et al. [19] presented the shrinking core (SC) model, which describes a process of irreversible desorption that is followed by diffusion through the pores of a porous solid. When the mass transfer rate of the solute in the non-extracted inner region is significantly slower than in the outer region, where most of the solute has already been extracted or when the solute concentration exceeds its solubility in the solvent, a distinct boundary may form between the inner and outer regions. As extraction progresses, the core of the inner region shrinks. The model envisions supercritical CO₂ extraction as a sharp, inward-moving front, with a completely non-extracted core ahead of the front and a fully extracted shell behind it.

Sovova [3] proposed the broken-and-intact cell (BIC) model, which assumes that a portion of the solute, initially stored within plant structures and protected by cell walls, is released during the mechanical breakdown of the material. The solute located in the region of broken cells near the particle surface is directly exposed to the solvent, while the core of the particle contains intact cells with undamaged walls. This model describes three extraction phases: a fast extraction phase for accessible oil, a transient phase, and a slow phase controlled by diffusion. The model has been successfully applied to the extraction of grape oil (Sovová et al. [20]) and caraway oil (Sovova et al. [21]).

The supercritical fluid extraction (SFE) process can be treated similarly to heat transfer, considering solid particles as hot balls cooling down in a uniform environment. Bartle et al. [22] introduced the hot ball diffusion (HBD) model, where spherical particles with a uniformly distributed solute diffuse similarly to heat diffusion. Unlike the BIC model, where the solute is readily available on the particle surface, the HBD model is suited for systems with small quantities of extractable materials and is not limited by solubility. The model is particularly relevant when internal diffusion controls mass transfer, allowing results from single particles to be extended to the entire bed under uniform conditions. Reverchon et al. [2] have further elaborated on the HBD model and used it to simulate extraction processes.

Reverchon [23] proposed a model for extraction of essential oils, which are mainly located inside the vegetable cells in organules called vacuoles. Only a small fraction of essential oil might be near the particle surface due to the breaking up of cells during grinding or in epidermal hairs located on the leaf surface. The fraction of oil freely available on the particle surface should not be significant in the case of SFE from leaves. Consequently, SFE of essential oil from leaves should be mainly controlled by internal mass-transfer resistance. Therefore, the external mass-transfer coefficient was neglected in the development of the model of Reverchon [23]. The mass balances were developed with the additional

hypotheses that axial dispersion can be neglected and that the solvent density and flow rate are constant along the bed.

This work builds upon the linear kinetic model suggested by Reverchon [23], deriving fundamental governing equations to develop a comprehensive model for the chamomile oil extraction process. This model aims at control-oriented simplicity, assuming semi-continuous operation within a cylindrical vessel. The process involves a supercritical solvent being pumped through a fixed bed of finely chopped biomass to extract the solute, followed by separation of the solvent and solute in a flush drum to collect the extract. Parameters such as pressure (P), feed flow rate (F) and inlet temperature (T_{in}) are adjustable and measurable, while the outlet temperature (T_{out}) and the amount of product at the outlet can only be monitored. Figure 1 presents a simplified process flow diagram.

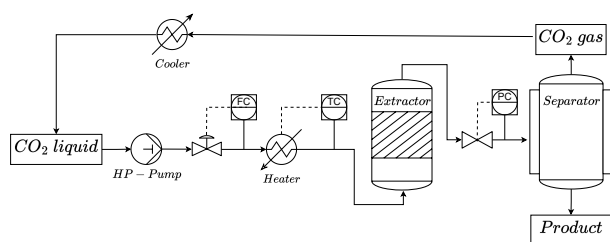


Figure 1: Process flow diagram.

The primary aim of this study is to analyse the influence of changes in operating conditions on the process model developed by Sliczniuk and Oinas [24]. Understanding how variations in parameters and control variables affect the model's behaviour is essential for improving its accuracy, robustness, and practical applicability. To achieve this, a comprehensive sensitivity analysis is employed, serving as a systematic approach to assess the impact of model parameters and controls on the outputs.

The results of this sensitivity analysis are valuable for multiple purposes. First, they help identify sources of uncertainty within the model, highlighting which parameters most significantly influence predictions. Second, they provide insights for potential model simplification, allowing the exclusion of non-essential parameters without compromising accuracy. Finally, the analysis can detect inconsistencies or errors by uncovering unexpected relationships between inputs and outputs, which may signal flaws in the model's assumptions or structure.

A range of sensitivity analysis methods can be utilized to achieve these objectives, including:

- One-at-a-time method
- Derivative-based local methods
- Variance-based methods

Different supercritical extraction models have been analysed using various sensitivity analysis techniques in the literature. For instance, Fiori et al. [25] performed sensitivity

calculations by varying parameters within their confidence intervals and observing the changes in model results. Their analysis revealed that the particle diameter and internal mass transfer coefficient significantly influence extraction during the diffusion-control regime.

Santos et al. [26] considered the model of Sovova [3] for semi-continuous isothermal and isobaric extraction processes using carbon dioxide as a solvent. They conducted a parametric sensitivity analysis using a two-level factorial design, disturbing model parameters by 10% and analysing their main effects. They proposed strategies for high-performance operation based on sensitivities related to superficial velocity, particle diameter, initial solute concentration in the solid phase and solute concentration in the fluid phase at the extractor inlet.

Hatami and Ciftci [27] performed a one-factor-at-a-time sensitivity analysis to assess the response of net present value (NPV) to variations in technical and economic variables. Their study consisted of two parts. The first part examined how net present value is influenced by changes in individual technical and economic parameters, keeping the extractor volume constant at 300 L. The second part investigated the effects of varying the extractor volume (from 1 to 600 L) on the project's profitability. The authors found that the raw material price, discount rate and residence time had the biggest impact on NPV.

The paper of Maly and Petzold [28] addresses the development of numerical methods and software tools for sensitivity analysis in differential-algebraic equation systems. The paper discusses the mathematical foundations of the local sensitivity analysis methods employed, providing a thorough analysis of their performance. The authors presented variety of problems and to prove the efficient of their method.

In his Turanyi [29] presents a comprehensive review of sensitivity analysis methods applied to spatially homogeneous, constant-parameter chemical kinetic systems. The work delves into various sensitivity analysis techniques, including concentration sensitivity, rate sensitivity, and feature sensitivity analysis. The paper explores practical applications of these methods in chemical kinetics, demonstrating their utility in understanding and predicting system behavior.

Fishtik and Geana [30] applied the local sensitivity analysis of an multiple chemical reaction system with several pure condensed phases and one ideal gas phase. It was shown that the system's response may be presented as a sum of contributions coming only from response equilibria defined by gas phase species. The effect of temperature, pressure and initial amounts of species is analyzed for gasification of coal.

Saltelli et al. [31] provide a comprehensive review of sensitivity analysis methods applicable to chemical modeling. The authors discuss both local and global techniques, emphasizing the importance of global sensitivity analysis in understanding complex chemical systems. The manuscript highlight the role of variance-based methods, in quantifying the influence of input parameters on model outputs.

Unlike previous studies, this work focuses on derivative-based local sensitivity analysis to evaluate the robustness and

predictive accuracy of a mathematical model for chamomile oil extraction under varying operational conditions. The analysis specifically examines the impact of operating condition changes, providing new insights into model development and optimizing extraction performance.

2. Materials and methods

2.1. Governing equations

The governing equations for a quasi-one-dimensional flow were derived following the work of Jr. [32]. A quasi-one-dimensional flow refers to a fluid flow scenario assuming that the flow properties are uniformly distributed across any cross-section. This simplification is typically applied when the flow channel's cross-sectional area changes, such as through irregular shapes or partial filling of an extractor. According to this assumption, velocity and other flow properties change solely in the flow direction.

As discussed by Jr. [33], all flows are compressible, but some of them can be treated as incompressible since the velocities are low. This assumption leads to the incompressible condition: $\nabla \cdot u = 0$, which is valid for constant density (strict incompressible) or varying density flow. The assumption allows for removing acoustic waves and large perturbations in density and/or temperature. In the 1-D case, the incompressibility condition becomes $\frac{du}{dz} = 0$, so the fluid velocity is constant along the z -direction.

The set of quasi-one-dimensional governing equations in Cartesian coordinates is described by Equations 1 - 3:

$$\frac{\partial(\rho_f A_f)}{\partial t} + \frac{\partial(\rho_f A_f v)}{\partial z} = 0, \quad (1)$$

$$\frac{\partial(\rho_f v A_f)}{\partial t} + \frac{\partial(\rho_f A_f v^2)}{\partial z} = -A_f \frac{\partial P}{\partial z}, \quad (2)$$

$$\frac{\partial(\rho_f e A_f)}{\partial t} + \frac{\partial(\rho_f A_f v e)}{\partial z} = -P \frac{\partial(A_f v)}{\partial z} + \frac{\partial}{\partial z} \left(k \frac{\partial T}{\partial z} \right), \quad (3)$$

where ρ_f is the density of the fluid, A_f is the function which describes a change in the cross-section, v is the velocity, P is the total pressure, e is the internal energy of the fluid, t is time and z is the spatial direction.

2.2. Extraction model

2.2.1. Continuity equation

The previously derived quasi-one-dimensional continuity equation (Equation 1) is redefined by incorporating the function $A_f = A\phi$. This modification distinguishes constant and varying terms, where the varying term accounts for changes in the cross-sectional area available for the fluid. Equation 4 shows the modified continuity equation:

$$\frac{\partial(\rho_f \phi)}{\partial t} + \frac{\partial(\rho_f v A \phi)}{\partial z} = 0, \quad (4)$$

where A is the total cross-section of the extractor and ϕ describes porosity along the extractor.

Assuming that the mass flow rate is constant in time, the temporal derivative becomes the mass flux F , and the spatial derivative can be integrated along z as

$$\int \frac{\partial(\rho_f v A \phi)}{\partial z} dz = F \rightarrow F = \rho_f v A \phi \quad (5)$$

To simplify the system dynamics, it is assumed that F is a control variable and affects the whole system instantaneously (due to $\nabla \cdot u = 0$), which allows finding the velocity profile that satisfies mass continuity based on F , ϕ and ρ_f :

$$v = \frac{F}{\rho_f A \phi} \quad (6)$$

Similarly, superficial velocity may be introduced:

$$u = v \phi = \frac{F}{\rho_f A} \quad (7)$$

The fluid density ρ_f can be obtained from the Peng-Robinson equation of state if the temperature and thermodynamic pressure are known along z . Variation in fluid density may occur due to pressure or inlet temperature changes. In a non-isothermal case, in Equations 6 and 7, ρ_f is considered the average fluid density along the extraction column.

2.2.2. Mass balance for the fluid phase

Equation 8 describes the movement of the solute in the system, which is constrained to the axial direction due to the quasi-one-dimensional assumption. Given that the solute concentration in the solvent is negligible, the fluid phase is described as pseudo-homogeneous, with properties identical to those of the solvent itself. It is also assumed that the thermodynamic pressure remains constant throughout the device. The analysis further simplifies the flow dynamics by disregarding the boundary layer near the extractor's inner wall. This leads to a uniform velocity profile across any cross-section perpendicular to the axial direction. Thus, the mass balance equation includes convection, diffusion and kinetic terms representing the fluid phase behaviour:

$$\frac{\partial c_f}{\partial t} + \frac{1}{\phi} \frac{\partial(c_f u)}{\partial z} = \frac{1 - \phi}{\phi} r_e + \frac{1}{\phi} \frac{\partial}{\partial z} \left(D_e^M \frac{\partial c_f}{\partial z} \right), \quad (8)$$

where c_f represents the solute concentration in the fluid phase, r_e is the mass transfer kinetic term and D_e^M is the axial dispersion coefficient.

2.2.3. Mass balance for the solid phase

As given by Equation 9, the solid phase is considered stationary, without convection and diffusion terms in the mass balance equation. Therefore, the only significant term in this equation is the kinetic term of Equation 10, which connects the solid and fluid phases. For simplicity, the extract is represented by a single pseudo-component:

$$\frac{\partial c_s}{\partial t} = \underbrace{r_e}_{\text{Kinetics}} \quad (9)$$

2.2.4. Kinetic term

As the solvent flows through the fixed bed, CO_2 molecules diffuse into the pores, adsorb on the inner surface and form a film due to solvent-solid matrix interactions. The dissolved solute diffuses into the bulk from the particle's core through the solid-fluid interface, the pore and the film. Figure 2 shows the mass transfer mechanism, where the mean solute concentration in the solid phase is denoted by c_s , and the equilibrium concentrations at the solid-fluid interface are

denoted by c_s^* and c_p^* for the solid and fluid phases, respectively. The concentration of the solutes in the fluid phase in the centre of the pore is denoted by c_p . As the solute diffuses through the pore, its concentration changes, reaching c_{pf} at the opening. Then, the solute diffuses through the film around the particle and reaches bulk concentration c_f . The two-film theory describes the solid-fluid interface inside the pore. The overall mass transfer coefficient can be determined from the relationship between the solute concentration in one phase and its equilibrium concentration.

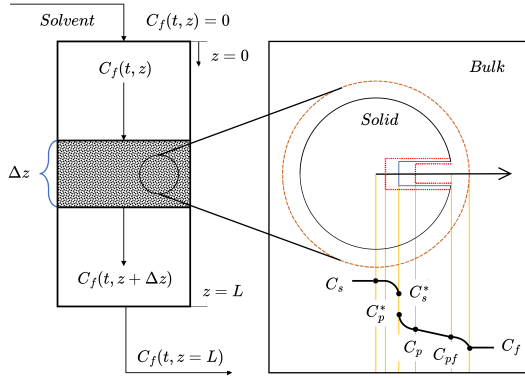


Figure 2: Mass transfer mechanism.

Bulley et al. [34] suggest a process where the driving force for extraction is given by the difference between the concentration of the solute in the bulk, c_f , and in the centre of the pore, c_p^* . The concentration c_p^* is in equilibrium with c_s according to the equilibrium relationship. The rate of extraction is thus $r_e (c_f - c_p^*(c_s))$. In contrast, Reverchon [23] proposes a driving force given by the difference between c_s and c_p^* . As given in Equation 10, the concentration c_p^* is determined by the equilibrium relationship with c_f :

$$r_e = \frac{D_i}{\mu l^2} (c_s - c_p^*), \quad (10)$$

where μ is sphericity, l a characteristic dimension of particles that can be defined as $l = r/3$, r is the mean particle radius, ρ_s is the solid density, D_i corresponds to the overall diffusion coefficient and c_p^* is the concentration at the solid-fluid interface (which, according to the internal resistance model, is supposed to be at equilibrium with the fluid phase).

According to Bulley et al. [34], a linear equilibrium relationship (Equation 11) can be used to find the equilibrium concentration of the solute in the fluid phase c_f^* based on the concentration of the solute in the solid phase c_s :

$$c_f^* = k_p c_s \quad (11)$$

The volumetric partition coefficient k_p acts as an equilibrium constant between the solute concentration in one phase and the corresponding equilibrium concentration at the solid-fluid interphase. Spiro and Kandiah [35] propose defining the mass partition coefficient k_m as

$$k_m = \frac{k_p \rho_s}{\rho_f} \quad (12)$$

According to Reverchon [23], the kinetic term becomes

$$r_e = -\frac{D_i}{\mu l^2} \left(c_s - \frac{\rho_s c_f}{k_m \rho_f} \right) \quad (13)$$

The choice between the two formulations should consider the system's properties and the underlying mass transfer assumptions. In systems where external mass transfer dominates, the resistance to solute transport primarily occurs at the fluid-solid interface, typically governed by the diffusion of solute across the fluid boundary layer. Under these conditions, the Bulley formulation is more appropriate, as it explicitly emphasizes the fluid-phase concentration (c_f) as the driving force for mass transfer through the term $c_f - c_p^*(c_s)$. Conversely, when internal mass transfer dominates, the resistance to solute transport is controlled by diffusion or adsorption processes within the solid phase. In such cases, the Reverchon formulation is more suitable, as it accounts for deviations in the equilibrium through the term $c_s - c_s^*(c_f)$.

2.2.5. Uneven distribution of solute in the solid phase

Following the idea of the broken-and-intact cell (BIC) model (Sovova [36]), the internal diffusion coefficient D_i is considered to be a product of the reference value of D_i^R and the exponential decay function γ , as given by Equation 14:

$$D_i = D_i^R \gamma(c_s) = D_i^R \exp \left(\gamma \left(1 - \frac{c_s}{c_{s0}} \right) \right), \quad (14)$$

where γ describes the curvature of the decay function. Equation 15 describes the final form of the kinetic term:

$$r_e = -\frac{D_i^R \gamma}{\mu l^2} \left(c_s - \frac{\rho_s c_f}{k_m \rho_f} \right) \quad (15)$$

The γ function limits the solute's availability in the solid phase. Similarly to the BIC model, the solute is assumed to be contained in the cells, some of which are open because the cell walls were broken by grinding, with the rest remaining intact. The diffusion of the solute from a particle's core takes more time than the diffusion of the solute close to the outer surface. The same idea can be represented by the decaying internal diffusion coefficient, where the decreasing term is a function of the solute concentration in the solid.

An alternative interpretation of the decay function γ involves considering the porous structure of the solid particles, where the pores are initially saturated with the solute. During extraction, the solute within these pores gradually dissolves into the surrounding fluid. Initially, the solute molecules near the pore openings dissolve and diffuse rapidly due to the short diffusion paths. As the extraction progresses, the dissolution front moves deeper into the pore structure, and solute from the inner regions of the pores begins to dissolve. The diffusion of solute molecules from the interior of the pores to the external fluid becomes progressively slower because the effective diffusion path length increases. This lengthening of the diffusion path enhances the mass transfer resistance, reducing the overall diffusion rate.

In an extreme case, this model could be compared with the shrinking core model presented by Goto et al. [19], where the particle radius decreases as the solute content in the solid

phase diminishes. In the SC model, the reduction in particle size leads to significant changes in both the diffusion path length and the surface area available for mass transfer. The diminishing particle size increases the diffusion path within the remaining solid core and decreases the external surface area, both of which contribute to a slower extraction rate. When comparing this to the varying diffusion coefficient in our model, some conceptual similarities can be noticed.

2.2.6. Empirical correlations

The empirical correlations for D_i and Y were derived by Sliczniuk and Oinas [24] and validated for temperatures of 30–40 °C, pressures of 100–200 bar, and mass flow rates of $3.33–6.67 \cdot 10^{-5}$ kg/s. Figures 3 and 4 show the results of multiple linear regression applied to solutions of parameter estimation and selected independent variables. The region marked with the white dashed line represents the confidence region, where the model has been tested. Both correlations should be equal or greater than zero to avoid unphysical behaviour such as reverse mass transfer. The multiple linear regression functions are combined with the rectifier function to ensure non-negativity.

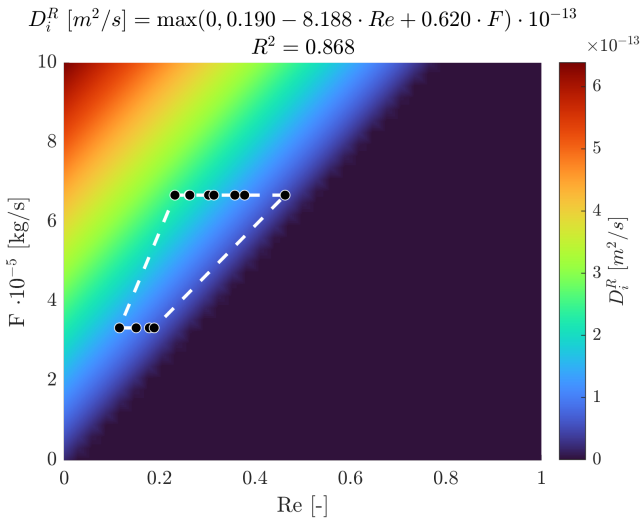


Figure 3: Multiple linear regression $D_i^R = f(Re, F)$.

2.2.7. Heat balance

The heat balance equation describes the evolution of the enthalpy in the system and is given by Equation 16:

$$\frac{\partial (\rho_f h A_f)}{\partial t} = -\frac{\partial (\rho_f h A_f v)}{\partial z} + \frac{\partial (P A_f)}{\partial t} + \frac{\partial}{\partial z} \left(k \frac{\partial T}{\partial z} \right) \quad (16)$$

If the value of enthalpy h is known from the time evolution of the energy equation, and pressure P is known from measurement, then temperature T can be reconstructed based on the departure function. The departure function is a mathematical function that characterizes the deviation of a thermodynamic property (enthalpy, entropy or internal energy) of a real substance from that of an ideal gas at the same temperature and pressure. The enthalpy departure

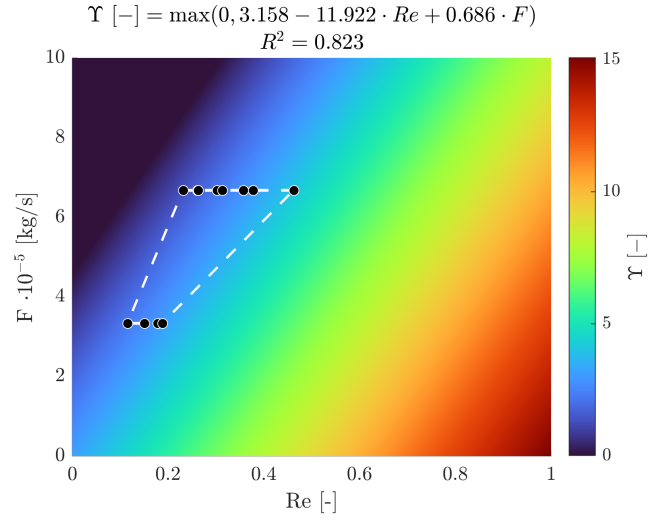


Figure 4: Multiple linear regression $Y = f(Re, F)$.

function, as presented by Gmehling et al. [37] for the Peng-Robinson equation of state, is defined by Equation 17:

$$h - h^{id} = RT \left[T_r(Z - 1) - 2.078(1 + \kappa)\sqrt{\alpha(T)} \ln \left(\frac{Z + (1 + \sqrt{2})B}{Z + (1 - \sqrt{2})B} \right) \right], \quad (17)$$

where α is defined as $(1 + \kappa(1 - \sqrt{T_r}))^2$, T_r is the reduced temperature, P_r is the reduced pressure, Z is the compressibility factor, κ is a quadratic function of the acentric factor and B is calculated as $0.07780 \frac{P_r}{T_r}$.

Equation 17 requires a reference state, which is assumed to be $T_{ref} = 298.15$ K and $P_{ref} = 1.01325$ bar.

A root finder can be used to find a temperature value which minimizes the difference between the value of enthalpy coming from the heat balance and the departure functions. The root-finding procedure has to be repeated at every time step to find a temperature profile along spatial direction z .

$$\min_T \left[\underbrace{h(t, x)}_{\text{Heat balance}} - \underbrace{h(T, P, \rho_f(T, P))}_{\text{Departure function}} \right]^2 \quad (18)$$

2.2.8. Pressure term

As explained in Section 2.1, this low velocity region pressure is nearly constant due to the small pressure wave propagation that occurs at the speed of sound. Under such conditions, the term $\partial P / \partial t$ can be approximated by a difference equation, describing the pressure change in the whole system. The pressure P in the system is considered a state variable, while the pressure in the new time-step P_{in} is considered a control variable.

$$\frac{\partial P}{\partial t} \approx \frac{P_{in} - P}{\Delta t} \quad (19)$$

Such a simplified equation allows for instantaneous pressure change in the system but does not consider a gradual

pressure build-up and the effects of pressure losses. In a real system, the dynamics of pressure change would depend on a pump and a back-pressure regulator.

2.2.9. Extraction yield

The process yield is calculated according to Equation 20, as presented by Sovova et al. [21]. The measurement equation evaluates the solute's mass at the extraction unit outlet and sums it up. The integral form of the measurement (Equation 20) can be transformed into the differential form (Equation 21) and augmented with the process model.

$$y = \int_{t_0}^{t_f} \frac{F}{\rho_f} c_f \Big|_{z=L} dt \quad (20)$$

$$\frac{dy}{dt} = \frac{F}{\rho_f} c_f \Big|_{z=L} \quad (21)$$

2.2.10. Initial and boundary conditions

It is assumed that the solvent is free of solute at the beginning of the process $c_{f0} = 0$, that all the solid particles have the same initial solute content c_{s0} and that the system is isothermal, hence the initial state is h_0 . The fluid at the inlet is considered not to contain any solute. The initial and boundary conditions are defined as follows:

$$\begin{aligned} c_f(t=0, z) &= 0 & c_s(t=0, z) &= c_{s0} & h(t=0, z) &= h_0 \\ c_f(t, z=0) &= 0 & h(t, z=0) &= h_{in} & \frac{\partial c_f(t, z=L)}{\partial x} &= 0 \\ \frac{\partial h(t, z=L)}{\partial x} &= 0 & c_s(t, z=\{0, L\}) &= 0 & y(0) &= 0 & P(0) &= P_0 \end{aligned}$$

2.2.11. Discretization methods

The method of lines is used to transform the process model equations into a set of ODEs denoted by $G(x; \Theta)$. The backward finite difference is used to approximate the first-order derivative, while the central difference scheme approximates the second-order derivative z direction. The length of the fixed bed is divided into N_z , i.e. equally distributed points in the z direction. The state-space model after discretization is denoted by x and defined as follows:

$$\dot{x} = \frac{dx}{dt} = \begin{bmatrix} \frac{dc_{f,1}}{dt} \\ \vdots \\ \frac{dc_{f,N_z}}{dt} \\ \frac{dc_{s,1}}{dt} \\ \vdots \\ \frac{dc_{s,N_z}}{dt} \\ \frac{dh_1}{dt} \\ \vdots \\ \frac{dh_{N_z}}{dt} \\ \frac{dP}{dt} \\ \frac{dy}{dt} \end{bmatrix} = \begin{bmatrix} G_1(c_f, c_s, h; \Theta) \\ \vdots \\ G_{N_z}(c_f, c_s, h; \Theta) \\ G_{N_z+1}(c_f, c_s, h; \Theta) \\ \vdots \\ G_{2N_z}(c_f, c_s, h; \Theta) \\ G_{2N_z+1}(c_f, c_s, h; \Theta) \\ \vdots \\ G_{3N_z}(c_f, c_s, h; \Theta) \\ G_{3N_z+1}(c_f, c_s, h; \Theta) \\ \underbrace{G_{3N_z+2}(c_f, c_s, h; \Theta)}_{G(x; \Theta)} \end{bmatrix}$$

where $x \in \mathbb{R}^{N_x=3N_z+2}$ and $\Theta \in \mathbb{R}^{N_\Theta=N_\theta+N_u}$, N_θ is the number of parameters and N_u is the number of control variables.

For a derivative to be conservative, it must form a telescoping series. In other words, only the boundary terms should remain after adding all terms coming from the discretization over a grid, and the artificial interior points should be cancelled out. Discretization is applied to the conservative form of the model to ensure mass conservation.

2.3. Local sensitivity analysis

Local derivative-based methods involve taking the total derivative of the state vector x with respect to the parameter space Θ . A set of derivatives, known as sensitivity equations, is integrated simultaneously with the process model. Sensitivity analysis shows how responsive the solution is to changes in the parameter Θ . As discussed by Dickinson and Gelinas [38], sensitivity analysis can be used to determine the influence of uncertainty on the solution of the original system. Another application of sensitivity analysis is to distinguish sensitive parameters from insensitive ones, which might be helpful for model reduction. Finally, from a control engineering point of view, sensitivity analysis allows sorting the control variables with respect to the level of effort required to change the model's output.

Following the work of Maly and Petzold [28], sensitivity equations can be defined as follows:

$$S(x; \Theta) = \frac{dx}{d\Theta} \quad (22)$$

The new system of equations can be obtained by taking the total derivative of S with respect to time t :

$$\dot{S}(x; \Theta) = \frac{dS(x; \Theta)}{dt} = \frac{d}{dt} \left(\frac{dx}{d\Theta} \right) = \frac{d}{d\Theta} \left(\frac{dx}{dt} \right) = \frac{dG(x; \Theta)}{d\Theta} \quad (23)$$

Sensitivity Equation 24 can be obtained by applying the definition of the total derivative to Equation 23:

$$\frac{dG(x; \Theta)}{d\Theta} = \underbrace{\frac{\partial G(x; \Theta)}{\partial x}}_{\bar{J}_x(x; \Theta)} \underbrace{\frac{\partial x}{\partial \Theta}}_{\bar{S}(x; \Theta)} + \underbrace{\frac{\partial G(x; \Theta)}{\partial \Theta}}_{\bar{J}_\Theta(x; \Theta)} \quad (24)$$

The Jacobian $\bar{J}_x(x; \Theta)$ represents the matrix of equations of size $N_x \times N_x$, where each equation $\bar{J}_x(n_x, n_x)$ is the derivative of $G_{n_x}(x; \Theta)$ with respect to the state variable x_{n_Θ} :

$$\bar{J}_x(x; \Theta) = \begin{pmatrix} \frac{\partial G_1(x; \Theta)}{\partial x_1} & \frac{\partial G_1(x; \Theta)}{\partial x_2} & \dots & \frac{\partial G_1(x; \Theta)}{\partial x_{N_x}} \\ \frac{\partial G_2(x; \Theta)}{\partial x_1} & \frac{\partial G_2(x; \Theta)}{\partial x_2} & \dots & \frac{\partial G_2(x; \Theta)}{\partial x_{N_x}} \\ \vdots & \vdots & \ddots & \vdots \\ \frac{\partial G_{N_x}(x; \Theta)}{\partial x_1} & \frac{\partial G_{N_x}(x; \Theta)}{\partial x_2} & \dots & \frac{\partial G_{N_x}(x; \Theta)}{\partial x_{N_x}} \end{pmatrix} \quad (25)$$

The sensitivity matrix $\bar{S}(x; \Theta)$ represents the matrix of equations of size $N_x \times N_\Theta$, where each entry $\bar{S}(n_x, n_\Theta)$ is the derivative of state variable x_{n_x} with respect to parameter Θ_{n_Θ} . Matrix $\bar{J}_x(x; \Theta)$ and $\bar{S}(x; \Theta)$ describe the indirect influence of Θ_{n_Θ} on the state space.

$$\bar{S}(x; \Theta) = \begin{pmatrix} \frac{\partial x_1}{\partial \Theta_1} & \frac{\partial x_1}{\partial \Theta_2} & \dots & \frac{\partial x_1}{\partial \Theta_{N_\Theta}} \\ \frac{\partial x_2}{\partial \Theta_1} & \frac{\partial x_2}{\partial \Theta_2} & \dots & \frac{\partial x_2}{\partial \Theta_{N_\Theta}} \\ \vdots & \vdots & \ddots & \vdots \\ \frac{\partial x_{N_x}}{\partial \Theta_1} & \frac{\partial x_{N_x}}{\partial \Theta_2} & \dots & \frac{\partial x_{N_x}}{\partial \Theta_{N_\Theta}} \end{pmatrix} \quad (26)$$

The Jacobian $\bar{J}_\Theta(x; \Theta)$ represents the matrix of equations of size $N_x \times N_\Theta$, where each sub-equation $\bar{J}_\Theta(n_x, n_\Theta)$ is the partial derivative of process model equation G_{n_x} with respect to parameter Θ_{n_Θ} . $\bar{J}_\Theta(n_x, n_\Theta)$ defines the direct effect of Θ_{n_Θ} on the state space:

$$\bar{J}_\Theta(x; \Theta) = \begin{pmatrix} \frac{\partial G_1(x; \Theta)}{\partial \Theta_1} & \frac{\partial G_1(x; \Theta)}{\partial \Theta_2} & \dots & \frac{\partial G_1(x; \Theta)}{\partial \Theta_{N_\Theta}} \\ \frac{\partial G_2(x; \Theta)}{\partial \Theta_1} & \frac{\partial G_2(x; \Theta)}{\partial \Theta_2} & \dots & \frac{\partial G_2(x; \Theta)}{\partial \Theta_{N_\Theta}} \\ \vdots & \vdots & \ddots & \vdots \\ \frac{\partial G_{N_x}(x; \Theta)}{\partial \Theta_1} & \frac{\partial G_{N_x}(x; \Theta)}{\partial \Theta_2} & \dots & \frac{\partial G_{N_x}(x; \Theta)}{\partial \Theta_{N_\Theta}} \end{pmatrix} \quad (27)$$

The augmented system containing the original set of equations $G(x; \Theta)$ and sensitivity equations is called $\mathbf{G}(x; \Theta)$. The size of $\mathbf{G}(x; \Theta)$ is equal to $N_s = N_x(N_\Theta + 1)$.

$$\mathbf{G}(x; \Theta) = \begin{bmatrix} G(x; \Theta) \\ \bar{J}_x(x; \Theta)\bar{S}(x; \Theta) + \bar{J}_\Theta(x; \Theta) \end{bmatrix} \quad (28)$$

The initial conditions are described as

$$\mathbf{G}(x(t_0); \Theta) = \begin{bmatrix} x(t_0), & \frac{dx(t_0)}{d\Theta_1}, & \dots, & \frac{dx(t_0)}{d\Theta_{N_\Theta}} \end{bmatrix}^T = \begin{bmatrix} x_0, & 0, & \dots, & 0 \end{bmatrix}^T \quad (29)$$

3. Results

The details of the process model and parameters are provided in Sliczniuk and Oinas [24]. The process model was validated in the following range of operating conditions: temperatures between 30 – 40°C, pressures between 100 – 200 bar, and mass flow rates between 3.33 – 6.67 · 10⁻⁵ kg/s. The sensitivity analysis is performed at the midpoint of the validated range: 35°C, 150 bar and 5 · 10⁻⁵ kg/s. This study examines the impact of pressure change on the state space and extraction yield through local sensitivity analysis. The local sensitivity analysis results can be interpreted as the system's response to an infinitesimal deviation in pressure.

As discussed in Section 2.1, at low velocity, the fluid can be treated as incompressible, which results in the instant propagation of pressure throughout the system, enabling a single pressure value to be considered for the entire system. In response to a pressure change, the energy equation experiences a simultaneous deviation across the entire spatial domain. This pressure change impacts the fluid temperature within the computational domain, while boundary values are constrained by conditions specified at the domain's extremes. Dirichlet boundary conditions impose a fixed temperature value at the inlet, creating a thermal gradient that propagates through the system. Alternatively, Neumann boundary conditions specify a heat flux at the extremes. The zero Neumann boundary conditions are applied to ensure a uniform response to the pressure change of temperatures at the inlet, outlet, and within the extractor.

Figure 5 illustrates the sensitivity of the solute concentration in the solid phase to pressure changes. As discussed in Section 2.2.1, the velocity of a fluid is inversely proportional to its density, indicating that increased pressure reduces the fluid velocity. This results in an extended residence time, allowing for longer interaction between the solute and solvent. Initially, the extraction process operates in the kinetic-controlled regime, where the concentration gradient is high, and solute solubility is the limiting factor. As noted by Sliczniuk and Oinas [24], the system is considered far from saturation, which explains the low initial system response. This low initial response was also observed in the sensitivity analysis by Fiori et al. [25]. The system's response becomes more pronounced as the concentration gradient decreases and the extraction shifts from the kinetic-controlled to the diffusion-controlled regime. In Figure 5, all $\frac{dc_s}{dP}$ values are negative, which indicates a faster mass transfer from the solid phase, corresponding to enhanced mass transfer. Over time, as the amount of solute decreases, it becomes a limiting factor, reducing the effect of pressure changes on the system. Eventually, the sensitivities approach zero asymptotically as the solute is washed out of the bed.

Figure 6 illustrates the sensitivity of solute concentration in the fluid phase to pressure changes. Compared to Figure 5, the dynamic behaviour of fluid phase sensitivities is more apparent. Due to advection, the sensitivities related to the fluid phase move across the system in the direction of flow. Initially, the system response is low despite the pressure increase improving mass transfer, reflecting the

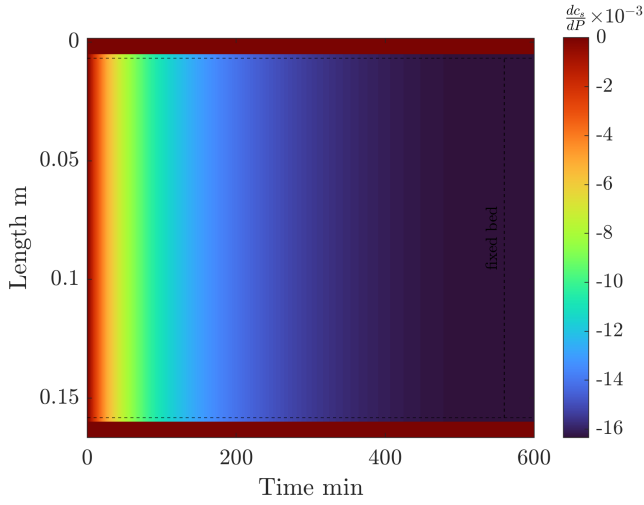


Figure 5: The effect of P_{in} change on C_s .

previously discussed idle period. As the process continues, the sensitivities increase, indicating faster mass transfer from the solid phase. The corresponding improvement in the mass transfer from the solid particles to the fluid is represented by the positive values in Figure 6. When the solute in the solid phase becomes a limiting factor, the extraction rate slows, and sensitivities eventually approach zero asymptotically.

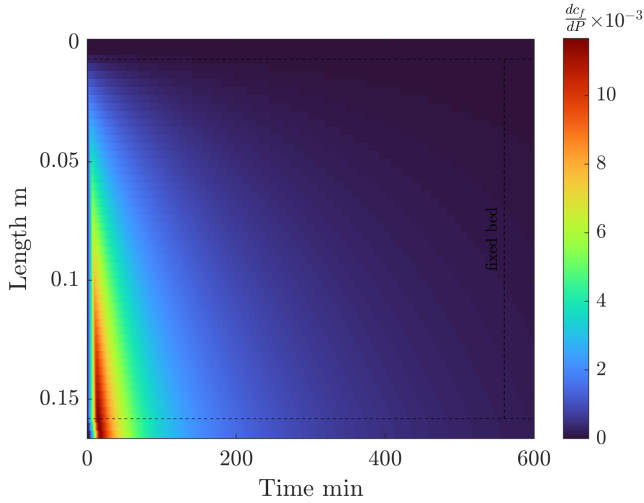


Figure 6: The effect of P_{in} change on C_f .

Figure 7 shows how sensitive the extraction yield is to pressure changes over an extended period for various pressure values. Initially, all sensitivity curves remain nearly flat, indicating a delayed response in the system. Due to the reduced fluid velocity, the solute reaches the extractor's outlet later, causing minor negative sensitivities to appear. Once the solute exits the extractor, the sensitivity curves rise. Positive yield sensitivities indicate improved process efficiency and enhanced mass transfer. The peak in $\frac{dy}{dP_{in}}$ represents the point of greatest deviation from the original

system. Eventually, the sensitivities decline and converge towards zero as the concentration gradient becomes a limiting factor, reducing the impact of enhanced mass transfer.

As presented in Figure 7, higher sensitivities are observed for a low-pressure system. At lower pressures, the supercritical fluid has lower density and solvating power, which results in less efficient extraction, as shown by data given by Sliczniuk and Oinas [24]. Small changes in pressure at low pressures can significantly impact the solute's solubility and, consequently, the extraction yield. Moreover, near the critical point, small changes in pressure can lead to significant changes in the physical properties of the supercritical fluid, such as density and viscosity, and therefore to higher sensitivity of the system state-space.

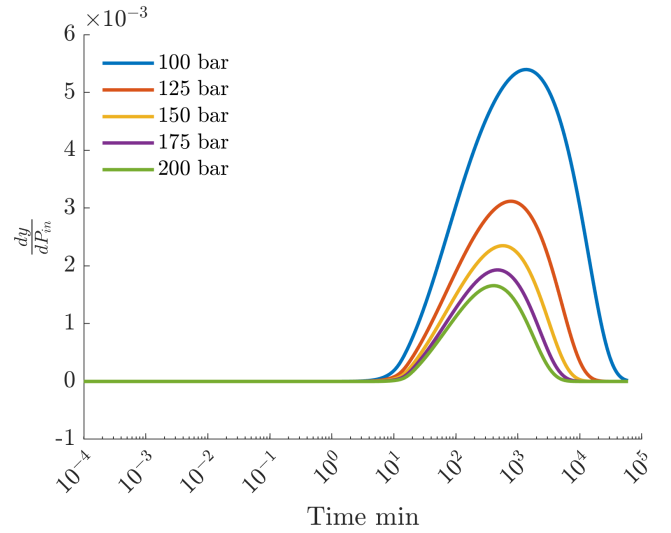


Figure 7: The effect of P_{in} change on $y(t)$.

One disadvantage of using local sensitivity coefficients is that they reflect the absolute changes, which might be difficult to compare among other models or scenarios. The normalized local sensitivity coefficients, defined as $\tilde{S}(x; \Theta) = \frac{dG(x; \Theta)}{d\Theta} \frac{\Theta}{G(x; \Theta)}$, which can be interpreted as the percentage change in the output per percentage change in the parameter, making it a unitless. Although, the interpretation is straightforward it should be noted that extrapolation for large change of the parameter might lead to incorrect results. In the case of dynamic systems, the outcome of local sensitivity analysis quantifies how the state of the system or its output varies over time with respect to small perturbations in parameters. If the perturbation is too large, then the definition of a derivative (or sensitivity equations) does not hold unless the system is linear. Consequently, the conclusions can not be scaled up to any size of parameter perturbation. Such a property of the local sensitivity analysis, makes it difficult to recreate in a laboratory conditions due to finite precision of machines.

Figure 8 illustrates the time evolution of the normalized sensitivity coefficients of the extraction yield. Unlike the raw sensitivities shown in Figure 7, the normalized sensitivities do not start from zero. This is due to the absence of the exact initial time point on the logarithmic time axis. The

pressure change impacts the system instantaneously and uniformly, leading to a rapid initial change in the normalized sensitivities. This behavior is analogous to a step change, where the system's response to the perturbation is immediate. Initially, the sensitivity curves remain flat. The increase in normalized sensitivity begins when the solute reaches the system's outlet and becomes detectable. After reaching their respective maxima, the sensitivity curves decrease as the concentration gradient driving the extraction diminishes over time. Ultimately, all curves approach zero, reflecting that all the solute was removed from the solid phase.

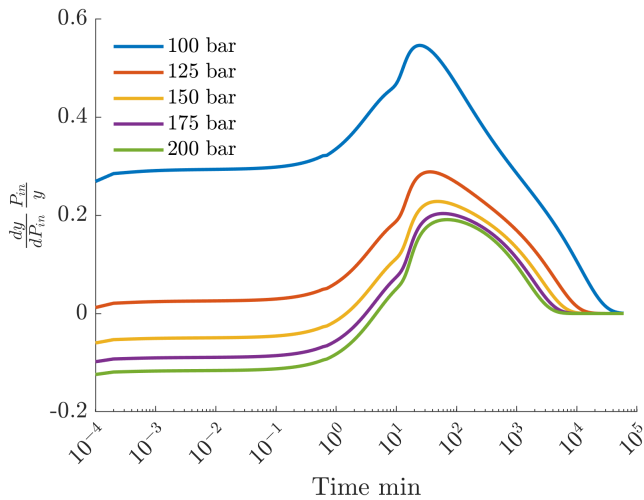


Figure 8: The effect of P_{in} change on $y(t)$ after normalization.

4. Conclusions

Sensitivity analysis is a tool used to understand how the parameters of the model affect its output. The presented formulation involves derivative-based local sensitivity analysis of the model solution with respect to selected parameters and controls. This work implemented automatic differentiation to derive the sensitivity equations. The results of the local sensitivity analysis and model specific and consider only a small region of parameter space, and the conclusions derived from such an analysis are limited to local conditions.

Although local sensitivity analysis can be performed with respect to any model parameter, this study focuses specifically on the effect of pressure. Under the selected operating conditions, it was observed for all the cases that a pressure increment enhances mass transfer, leading to a faster mass transfer of solute from the particles. Consequently, negative sensitivities can be observed in the diagram for the solid phase. The corresponding response in the fluid phase is characterized by positive-valued sensitivities, indicating that a greater amount of solute is transported into the fluid phase. As a result, the extraction yield is improved and characterized by positive sensitivities. Given that the results of the local sensitivity analysis depend on the operating point, the analysis was repeated at different pressures.

The normalized sensitivity coefficients provide a basis for generalizing the discussion of the results and comparing the influence of various parameters. The normalized sensitivity results were found to vary within the range of -0.17 to 0.6. The overall effect of each parameter can be evaluated by integrating the sensitivity curves. Similar to the analysis of yield sensitivity, the integration was performed using an extended simulation until all sensitivities approached zero. The cumulative integral values at pressures of 100, 125, 150, 175, and 200 bars are 31487, 8548, 4462, 2825, and 1939, respectively. The positive sign of these values indicates that increased pressure enhances the extraction yield under all investigated operating conditions. The order of magnitude of the integrals allows to identify at which operating conditions the pressure increase is the most influential.

It can be concluded that the system responses at low pressures were stronger than those at higher pressures. This behaviour can be explained by the rapid changes in solvent properties occurring near the critical point of CO_2 .

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897	Nomenclature		R_e	Reynolds number	-	933
898			r_e	Mass transfer kinetic term	kg/m ³ /s	934
899	\bar{J}_Θ	Jacobian matrix with respect to the parameters	S	Sensitivity equations		935
900	\bar{J}_x	Jacobian matrix with respect to the state variables	T	Temperature	K	936
901	\bar{S}	Sensitivity matrix	t	Time	s	937
902	\dot{S}	Time derivative of the sensitivity equations	t_0	Initial extraction time	s	938
903	G	Augmented system	t_f	Total extraction time	s	939
904	A	Total cross-section of the bed	T_{in}	Inlet temperature	K	940
905	A_f	Cross-section of the bed occupied by the fluid	T_{out}	Outlet temperature	K	941
906	c_f	Concentration of solute in the fluid phase	u	Superficial velocity	m/s	942
907	c_f^*	Concentration of solute at the solid-fluid interface	v	Linear velocity	m/s	943
908		kg/m ³	x	State vector		944
909	c_p	Concentration of solute in the core of a pore	y	Extraction yield	g	945
910	c_s	Concentration of solute in the solid phase	z	Spatial direction	m	946
911	c_s^*	Concentration of solute at the solid-fluid interface	Greek symbols			947
912		kg ³	ϵ	Unobservable error	g	948
913	c_{f0}	Initial concentration of solute in the fluid phase	γ	Decay function	-	949
914		kg/m ³	μ	Sphericity coefficient	-	950
915	c_{pf}	Concentration of solute at the pore opening	Φ	Bed porosity	-	951
916	c_{s0}	Initial concentration of solute in the solid phase	ρ_f	Fluid density	kg/m ³	952
917		kg/m ³	ρ_s	Bulk density of the solid	kg/m ³	953
918	D_e^M	Axial diffusion coefficient	σ	Standard deviation	g	954
919	D_i	Internal diffusion coefficient	Θ	Parameter space		955
920	D_i^R	Reference value of internal diffusion coefficient	θ	Vector of unknown parameters		956
921		m ² /s	Υ	Decay coefficient	-	957
922	e	Internal energy	Abberivations			958
923	F	Mass flow rate	BIC	Broke-and-Intact Cell model		959
924	G	Vector of discretized differential equations	HBD	Hot Ball Diffusion		960
925	h	Enthalpy	SC	Shrinking Core		961
926	k_m	Mass partition coefficient	SFE	Supercritical Fluid Extraction		962
927	k_p	Volumetric partition coefficient				
928	L	Length of the fixed bed				
929	l	Characteristic dimension of particles				
930	P	Pressure				
931	p	Probability distribution model				
932	r	Particle radius				