On a multi-species Cahn–Hilliard–Darcy tumor growth model with singular potentials

Sergio Frigeri * I

Kei Fong Lam[†] Elis

Elisabetta Rocca[‡]

Giulio Schimperna[‡]

September 6, 2017

Abstract

We consider a model describing the evolution of a tumor inside a host tissue in terms of the parameters φ_p , φ_d (proliferating and dead cells, respectively), \boldsymbol{u} (cell velocity) and n (nutrient concentration). The variables φ_p , φ_d satisfy a Cahn-Hilliard type system with nonzero forcing term (implying that their spatial means are not conserved in time), whereas \boldsymbol{u} obeys a form of the Darcy law and n satisfies a quasistatic diffusion equation. The main novelty of the present work stands in the fact that we are able to consider a configuration potential of singular type implying that the concentration vector (φ_p, φ_d) is constrained to remain in the range of physically admissible values. On the other hand, in view of the presence of nonzero forcing terms, this choice gives rise to a number of mathematical difficulties, especially related to the control of the mean values of φ_p and φ_d . For the resulting mathematical problem, by imposing suitable initial-boundary conditions, our main result concerns the existence of weak solutions in a proper regularity class.

Key words. Tumor growth, nonlinear evolutionary system, Cahn–Hilliard–Darcy system, existence of weak solutions, logarithmic potentials.
AMS subject classification. 35D30, 35Q35, 35Q92, 35K57, 76S05, 92C17, 92B05.

1 Introduction

Tumor growth remains an active area of scientific research due to the impact on the quality of life for those diagnosed with cancer. Starting with the seminal work of Burton [8] and Greenspan [33], many mathematical models have been proposed to emulate the complex biological and chemical processes that occur in tumor growth with the aim of better understanding and ultimately controlling the the behavior of cancer cells. In recent years, there has been a growing interest in the mathematical modelling of cancer, see for example [1, 3, 7, 19, 21, 23]. Mathematical models for tumor growth may have different analytical features: in the present work we are focusing on the subclass of continuum models, namely diffuse interface models. In this framework, the tumor and surrounding host tissue occupy regions of a domain and are subject to various balance laws mimicking the biological processes one would like to model. While it is intuitive to represent the

^{*}Dipartimento di Matematica e Fisica, Università Cattolica del Sacro Cuore, Via dei Musei 41, I-25121 Brescia, Italy (sergio.frigeri.sf@gmail.com).

[†]Department of Mathematics, The Chinese University of Hong Kong, Shatin, N.T., Hong Kong (kflam@math.cuhk.edu.hk).

[‡]Dipartimento di Matematica, Università di Pavia, and IMATI - C.N.R., Pavia, Via Ferrata 1, I-27100, Pavia, Italy (elisabetta.rocca@unipv.it, giusch04@unipv.it).

interfaces between the tumor and healthy tissues as idealized surfaces of zero thickness, leading to a sharp interface description that differentiates the tumor and the surrounding host tissue cell-by-cell, these kinds of sharp interface models are often difficult to analyze mathematically, and may break down when the interface undergoes a topological change. In particular, sharp interface models may fail to describe the process of metastasis, which is the spreading of cancer to other parts of the body, and the primary characteristic that makes cancer so deadly.

On the other hand, diffuse interface models consider the interface between the tumor and the healthy tissues as a narrow layer in which tumor and healthy cells are mixed. This alternative representation of the interface gives rise to model equations that are better amenable to mathematical analysis, and the mathematical description remains valid even when the tumor undergoes topological changes. Hence, the recent efforts in the mathematical modeling of tumor growth have been mostly focused on diffuse interface models, see for example [18, 19, 22, 29, 31, 34, 40, 44], and their numerical simulations demonstrating complex changes in tumor morphologies due to mechanical stresses and interactions with chemical species such as nutrients or toxic agents.

The interaction of multiple tumor cell species can be described by using multiphase mixture models (see, e.g., [1, 20, 22, 29, 42, 44]). Indeed, using multiphase porous media mechanics, the authors of [42] represented a growing tumor as a multiphase medium containing an extracellular matrix, tumor and host cells, and interstitial liquid. Numerical simulations were also performed that characterized the process of cancer growth in terms of the initial tumor-to-healthy cell density ratio, nutrient concentration, mechanical strain, cell adhesion, and geometry. The interactions of a growing tumor and a basement membrane were studied in [5]. In [9] the authors adapted the approach from [5] to the case of multiphase mixture models. Although the model studied in [9] contains numerous simplifications, the underlying approach has been successfully used by other authors, also with the purpose of comparing the outcome of numerical tests with experimental data (e.g., [22]). Furthermore, the modeling strategy and the numerical methods presented in [9] are generalizable to more sophisticated and thermodynamically consistent models that account for additional biophysical details (e.g., [19]). For these reasons, we will use here a modeling approach closely related to that of [9], with few modifications that will be detailed here below.

In terms of the theoretical analysis of diffuse interface models, most of the recent literature is restricted to the two-phase variant, i.e., models that only account for the evolution of a tumor surrounded by healthy tissue. In this setting, there is no differentiation among the tumor cells that exhibit heterogeneous growth behavior, and consequently this kind of two-phase models are just capable to describe the growth of a young tumor before the onset of quiescence and necrosis. Analytical results related to existence of weak and strong solutions, and, in some cases, continuous dependence on data, asymptotic limits and longtime behavior have been established in [11, 13, 15, 24, 25, 26, 27, 28, 41] for tumor growth models based on the coupling of Cahn–Hilliard (for the tumor density) and reaction– diffusion (for the nutrient or other chemical factors) equations, and in [27, 32, 35, 37, 38] for models of Cahn–Hilliard–Darcy type. There have also been some studies involving the optimal control and sliding modes for diffuse interface tumor growth, see, e.g., [12, 14, 30].

Comparatively, there have been fewer analytical results for the multi-phase variants, which distinguish between the proliferating and necrotic tumor cells. We just mention and briefly discuss the recent works of [20, 29]. In [29], a Cahn–Hilliard–Darcy model is derived to describe multiphase tumor growth taking interactions with multiple chemical species into account as well as the simultaneous occurrence of proliferating, quiescent

and necrotic regions. A new feature of the modeling approach of [29] is that a volumeaveraged velocity is used, which dramatically simplifies the resulting equation for the mixture velocity. With the help of formally matched asymptotic analysis the authors also develop new sharp interface models. On the other hand, in [20], the authors analyze the multi-species Cahn-Hilliard-Darcy tumor model of [9] in the case of constant and identical mobilities for all the species and establish the existence of a weak solution. We actually believe it is important to analyze these multi-species models, as once the tumor experiences quiescence and necrosis due to lack of nutrient, it will start secreting growth factors that induce the development of new blood vessels towards the tumor, a process known as angiogenesis, and it is through these new blood vessels that tumors tend to metastasize into other parts of the body.

In this contribution, we actually consider a multi-species tumor model posed in a smooth bounded domain $\Omega \subset \mathbb{R}^d$, $d \in \{2, 3\}$, and over a reference time interval (0, T) with no restriction on the magnitude of T. Our model describes the evolution of proliferating tumor cells, necrotic tumor cells and healthy host cells. We denote the corresponding volume fractions as $\varphi_p, \varphi_d, \varphi_h \in [0, 1]$, respectively, so that $\varphi_p + \varphi_d + \varphi_h = 1$ almost everywhere in $\Omega \times (0, T)$. By this relation, it suffices to track the evolution of φ_p and φ_d in order to deduce the evolution of φ_h ; for this reason it is also natural to assume that the vector $\boldsymbol{\varphi} := (\varphi_p, \varphi_d)^{\top}$ lies in the simplex $\Delta := \{\boldsymbol{y} \in \mathbb{R}^2 : 0 \leq y_1, y_2, y_1 + y_2 \leq 1\} \subset \mathbb{R}^2$. This constraint will be one of the key points in our approach and we will explain below how it is enforced by the equations.

In order to introduce our mathematical model we start by making some assumptions on the biological mechanisms experienced by the tumor cells. Let n denote the concentration of a nutrient chemical species that is present in Ω . Then, the proliferating tumor cells may grow by consuming the nutrient, and may die by apoptosis or necrosis in case of nutrient deficiency. Correspondingly, necrotic cells will increase in mass due to the apoptosis/necrosis of proliferating tumor cells, but they may also disintegrate back into basic components. On the other hand, the dynamics of the healthy cells occurs on a much slower timescale than that of the tumor cells, since we expect that compared to the strictly regulated mitosis cycle of the healthy cells, in the tumor certain growth-inhibited proteins have been switched off by mutations and leads to unregulated growth behavior which is only limited by the supply of nutrients.

Assuming the cells are tightly packed and move together, we may introduce a cell velocity field \boldsymbol{u} which is the same for all types of cells. Letting \boldsymbol{J}_i , $i \in \{p, d, h\}$, denote the mass fluxes for the tumor cells, then the general balance law for the volume fractions reads as

$$\partial_t \varphi_i + \operatorname{div}(\varphi_i \boldsymbol{u}) = -\operatorname{div} \boldsymbol{J}_i + S_i \text{ for } i \in \{p, d, h\},$$
(1.1)

where, in view of the above discussion, we set $S_h = 0$, whereas S_p , S_d may depend on n, φ_p and φ_d . Note that it is sufficient to determine the evolution of φ_p and φ_d in view of the fact that φ_h can be determined at any point as $1 - \varphi_p - \varphi_d$. We assume that the tumor growth process tends to evolve towards (local) minima of the following free energy functional $E(\varphi_p, \varphi_d)$ of Ginzburg–Landau type:

$$E(\varphi_p, \varphi_d) := \int_{\Omega} F(\varphi_p, \varphi_d) + \frac{1}{2} \left| \nabla \varphi_p \right|^2 + \frac{1}{2} \left| \nabla \varphi_d \right|^2 dx,$$
(1.2)

where F is a multi-well configuration potential for the variables φ_p and φ_d .

More precisely, we assume F to be the sum of a smooth non-convex part F_1 and of a nonsmooth singular convex part F_0 . Namely, F_0 is identically set to $+\infty$ outside the set

 $\overline{\Delta} = \{(s,r) \in \mathbb{R}^2 : s \ge 0, r \ge 0, s + r \le 1\} \text{ of the "physically admissible" configurations.}$ In other words, if $\int_{\Omega} F_0(\varphi_p, \varphi_d) dx$ is finite, then we necessarily have $\varphi_p, \varphi_d \ge 0$ and $\varphi_p + \varphi_d \le 1$. Moreover, as a further consequence, due to the fact that $\varphi_h = 1 - \varphi_p - \varphi_d$, we also have $0 \le \varphi_h \le 1$. In other words, the finiteness of the "configuration energy" $\int_{\Omega} F(\varphi_p, \varphi_d) dx$ automatically ensures the natural bounds

$$0 \le \varphi_p, \varphi_d, \varphi_h \le 1 \text{ a.e. in } \Omega.$$
 (1.3)

An example of singular potential F_0 that we can include in our analysis and that was already introduced in [44] is

$$F_0(\varphi_p, \varphi_d) := \varphi_p \log \varphi_p + \varphi_d \log \varphi_d + (1 - \varphi_p - \varphi_d) \log(1 - \varphi_p - \varphi_d).$$
(1.4)

This can be seen as a generalization of the standard one-component logarithmic potential commonly used in the framework of Cahn-Hilliard equations (cf. for example [10, 39]). Let us also comment that, in light of the above considerations, the pure phase consisting of proliferating tumor cells is characterized by the region { $\varphi_p = 1, \ \varphi_d = \varphi_h = 0$ }, whereas the pure phase corresponding to the necrotic cells is the region { $\varphi_d = 1, \ \varphi_p = \varphi_h = 0$ }. The fluxes $J_i, \ i = p, d$, are defined as follows:

$$J_i = -M_i \nabla \mu_i, \quad \mu_i := \frac{\delta E}{\delta \varphi_i} = -\Delta \varphi_i + F_{,\varphi_i} \text{ for } i = p, d,$$

where M_p, M_d are positive constants which can be different from each other. We set $J_h = -J_p - J_d$, then upon summing up the three equations (1.1), for i = p, d, h, using the fact that $\varphi_p + \varphi_d + \varphi_h = 1$ and $S_h = 0$, we deduce the following relation:

$$\operatorname{div} \boldsymbol{u} = S_p + S_d =: S_t \,.$$

The velocity field \boldsymbol{u} is assumed to fulfill Darcy's law (cf. [44]):

$$\boldsymbol{u} = -\nabla q - \varphi_p \nabla \mu_p - \varphi_d \nabla \mu_d,$$

where q denotes the cell-to-cell pressure and the subsequent two terms have the meaning of Korteweg forces. Regarding the nutrient n, since the time scale of nutrient diffusion is much faster (minutes) than the rate of cell proliferation (days), the nutrient is assumed to evolve quasi-statically:

$$0 = -\Delta n + \varphi_p n, \tag{1.5}$$

where the second term on the right-hand side models consumption by the proliferating tumor cells. Notice that we have neglected the nutrient uptake by host tissue because this is small compared with the uptake by tumor cells (e.g., see [44] for more details). Moreover we have also neglected here the capillarity term $T_C = B(\varphi_p, \varphi_d)(n_C - n)$ considered, e.g., in [20], just for the sake of simplicity. Actually, we could include it in our analysis assuming that the function $B(\varphi_p, \varphi_d)$ is smooth and non-negative and the coefficient n_C is strictly between 0 and 1 (cf, e.g., [20, (2.6)]). These assumptions would indeed guarantee the maximum principle is still true for the modified nutrient equation. Hence, setting $Q := \Omega \times (0, T)$ and $\Gamma := \partial \Omega \times (0, T)$, we are led to the following multi-species tumor model:

$$\partial_t \varphi_p = M_p \Delta \mu_p - \operatorname{div}(\varphi_p \boldsymbol{u}) + S_p, \quad \mu_p = F_{\varphi_p} - \Delta \varphi_p \text{ in } Q,$$
 (1.6a)

$$\partial_t \varphi_d = M_d \Delta \mu_d - \operatorname{div}(\varphi_d \boldsymbol{u}) + S_d, \quad \mu_d = F_{,\varphi_d} - \Delta \varphi_d \text{ in } Q,$$
 (1.6b)

$$S_p = \Sigma_p(n, \varphi_p, \varphi_d) + m_{pp}\varphi_p + m_{pd}\varphi_d \qquad \text{in } Q, \qquad (1.6c)$$

$$S_d = \Sigma_d(n, \varphi_p, \varphi_d) + m_{dp}\varphi_p + m_{dd}\varphi_d \qquad \text{in } Q, \qquad (1.6d)$$

$$\operatorname{div} \boldsymbol{u} = S_p + S_d \qquad \qquad \text{in } Q, \qquad (1.6e)$$

$$\boldsymbol{u} = -\nabla q - \varphi_p \nabla \mu_p - \varphi_d \nabla \mu_d \qquad \text{in } Q, \qquad (1.6f)$$

$$0 = -\Delta n + \varphi_p n \qquad \qquad \text{in } Q, \qquad (1.6g)$$

where the precise assumptions on the mass source terms Σ_p , Σ_d will be specified in Section 2.

We just give here an example of source terms in (1.6) that fulfill the assumption (2.2) stated in Section 2. Namely, we may set

$$S_p = \lambda_M g(n) - \lambda_A \varphi_p, \qquad (1.7)$$

$$S_d = \lambda_A \varphi_p - \lambda_L \varphi_d, \tag{1.8}$$

for positive constants λ_M , λ_A , λ_L and a bounded positive function g such that $0 < g(s) \leq 1$. The archetypal example is $g(s) = \max(n_c, \min(s, 1))$ for some constant $n_c \in (0, 1)$. The biological effects that we want to model here are: the growth of the proliferating tumor cells due to nutrient consumption at a constant rate λ_M , the death of proliferating tumor cells at a constant rate λ_A , which leads to a source term for the necrotic cells, and the lysing/disintegration of necrotic cells at a constant rate λ_L .

We supplement the system with the following boundary conditions:

$$M_i \partial_{\boldsymbol{n}} \mu_i - \varphi_i \boldsymbol{u} \cdot \boldsymbol{n} = 0, \quad \partial_{\boldsymbol{n}} \varphi_i = 0, \quad q = 0, \quad n = 1 \text{ on } \Gamma,$$
(1.9)

where ∂_n denotes the outer normal derivative to $\Gamma = \partial \Omega$, and with the initial conditions

$$\varphi_p(x,0) = \varphi_{p,0}(x), \quad \varphi_d(x,0) = \varphi_{d,0}(x) \text{ in } \Omega.$$
(1.10)

Note that, implicitly, we are also assuming that $\varphi_h(x,0) = 1 - \varphi_{p,0}(x) - \varphi_{d,0}(x)$ in Ω .

Let us now compare (1.6) and the model of [9] which was analyzed by [20]:

- We use a simpler nutrient equation (1.6g) which only accounts for diffusion and consumption by the proliferating tumor cells. In particular, we have neglected the nutrient capillary source term for simplicity (though, as remarked before, such a term could be treated by standard modifications).
- In [9], the effect of a basement membrane on the growing tumor is also considered, which leads to additional coupling of the model with a Cahn-Hilliard equation transported by the velocity u. In this work we do not consider such effects.
- The key distinction is that in our choice of a multi-well potential F in (1.2), we included interfacial energy for the proliferating-necrotic tumor interface and also for the tumor-host interfaces. On the other hand, in [9] the free energy depends only on the total tumor volume fraction $\varphi_T = \varphi_p + \varphi_d$, i.e., $E(\varphi_T) = \int_{\Omega} f(\varphi_T) + \frac{1}{2} |\nabla \varphi_T|^2 dx$ for scalar double-well potential f with minima at 0 and 1. This reduction to the total tumor volume fraction implies that the proliferating-necrotic tumor interface in [9] is not energetic.

- More precisely, in [9], like in the multiphase models studied in [22, 44, 45], the differentiation between proliferating and necrotic tumor cells is done a posteriori based on the local density of nutrients after computing φ_T . In contrast, our model (1.6) follows a similar approach to [29] in which φ_p and φ_d are computed without any post processing.
- Moreover, we consider here different boundary conditions with respect to [9], where a zero Dirichlet boundary datum was taken for the chemical potentials, while here we consider a coupled condition for μ_i and \boldsymbol{u} (the first of (1.9)). It is worth noting that the (easier) case of Dirichlet boundary conditions for μ_i could also be treated, but we preferred to handle (1.9) which seems to be more reasonable from the modeling point of view. On the contrary, the case of no-flux conditions for μ_i (which would also be meaningful) seems not easy to be treated mathematically.
- Finally, differently from [20], we assume here that different cell species are characterized by different mobility coefficients. Actually, this choice gives rise to a number of mathematical complications. In particular, here we cannot reduce the evolution of the tumor cells (as was done in [20]) to a single Cahn–Hilliard equation coupled with a transport-type relation, but we need to consider a vectorial Cahn–Hilliard system, for which, however, several issues are still open. On the other hand, here we get stronger regularity due to the fact that we do not have a transport equation anymore.

Mathematically speaking, the main novelty of our model, and also its main difficulty from the analytical point of view, comes from the singular component F_0 of the configuration potential coupled with the nonzero source terms in the Cahn-Hilliard relations (1.6a)-(1.6b). Indeed, integrating the first relations in (1.6a), (1.6b) we obtain an evolution law for the spatial mean values $y_i := \frac{1}{|\Omega|} \int_{\Omega} \varphi_i dx$ of φ_i for i = p, d (cf. (5.2) below) which is satisfied by any hypothetical solution to the system. Such a relation, however, does not involve directly the singular part F_0 . Hence, the evolution of y_p, y_d are not automatically compatible with the physical constraint (1.3) and this compatibility (i.e., the fact that y_p , y_d remain well inside the set of meaningful values) has to be carefully proved (see Subsec. 5.1) by assuming proper conditions on coefficients and making a careful choice of the boundary conditions. In particular, the first condition in (1.9) linking the boundary values of $\boldsymbol{u}, \varphi_i$ and μ_i seems to be necessary in order for our arguments to work. It is worth noting that very few results are nowadays available for multi-component Cahn-Hilliard systems. Among these, we mention the recent contributions [4, 17, 36], related to the case of regular (multi-well) potentials, whereas up to our knowledge, the only paper dealing with a singular multi-well potential like the logarithmic one (1.4) introduced in [44] is [16]. Indeed, some estimates proved in [16] will be used in the proof of our results.

Concerning the approach we employed to prove existence of weak solutions to system (1.6a)-(1.6g), the first step is to consider a regularized version of this problem, which is obtained by replacing the singular potential F_0 by a regular one depending on an approximation parameter $\varepsilon > 0$, and also by introducing some suitable truncation functions. We then present two independent methods that permit us to prove existence of a solution to the regularized system. The first one is based on a further regularization and a Schauder fixed point argument, whereas the second proof exploits a direct implementation of a Faedo-Galerkin scheme. Actually, we have decided to detail two different proofs, since, in our opinion, both of them present some independent interest. Indeed, the first method only exploits elementary existence and uniqueness results methods for PDEs and also permits us to split our complicated model into its basic components and show how

they can be treated separately. On the other hand, the Faedo-Galerkin method is more direct (no further regularizing terms are introduced), and constructive (hence, it may be used for a numerical approximation of the problem).

Plan of the paper. The assumptions and main results are stated in Section 2. The proof is carried out in the remainder of the paper and is subdivided into several steps: namely, in Section 3 regularized version of our model is introduced, where the singular part of the potential F_0 is replaced by a smooth approximation and some terms are truncated in order to maintain some boundedness properties that are used in the a priori estimates. Moreover, still in Section 3, the fixed point method for existence is outlined, whereas the alternative Faedo-Galerkin procedure is presented in Section 4. Once existence is established for the regularized model, in Section 5 some bounds that are independent of the regularization parameters are derived in order to pass to the limit in the approximation scheme via compactness tools and to obtain in this way existence of weak solutions for the original system.

2 Assumptions and main result

We start presenting our assumptions on parameters and data:

Assumption 2.1. (A1) M_p, M_d are strictly positive constants.

(A2) We set $\Sigma := (\Sigma_p, \Sigma_d)$ and denote as $\underline{M} = (m_{ij}), i, j \in \{p, d\}$, the matrix of the coefficients in (1.6c), (1.6d). Then we assume that there exists $c \ge 0$ such that

$$\Sigma \in C^{0,1}(\mathbb{R}^3; \mathbb{R}^2), \quad \|\Sigma\|_{L^{\infty}(\mathbb{R}^3; \mathbb{R}^2)} + \|D\Sigma\|_{L^{\infty}(\mathbb{R}^3; \mathbb{R}^6)} \le c.$$
(2.1)

In other words, Σ is globally Lipschitz. More precisely, we assume that there exist a closed and sufficiently regular subset Δ_0 contained in the open simplex Δ and constants $K_{p,-}, K_{p,+}, K_{d,-}, K_{d,+} \in \mathbb{R}$, with $K_{p,-} \leq K_{p,+}$ and $K_{d,-} \leq K_{d,+}$, such that $\Sigma(\mathbb{R}^3) \subset [K_{p,-}, K_{p,+}] \times [K_{d,-}, K_{d,+}]$. Moreover, for any $\mathbf{x} = (x_p, x_d) \in [K_{p,-}, K_{p,+}] \times [K_{d,-}, K_{d,+}]$, there holds

$$(\underline{M}\boldsymbol{y} + \boldsymbol{x}) \cdot \boldsymbol{n} < 0 \text{ for all } \boldsymbol{y} \in \partial \Delta_0,$$
(2.2)

where **n** denotes the outer unit normal vector to Δ_0 .

(A3) The potential F is the sum of a convex part F_0 and of a (possibly nonconvex) perturbation F_1 . More precisely, we assume that $F_0 : \mathbb{R}^2 \to [0, +\infty]$, with the effective domain of F_0 (i.e., the set where F_0 assumes finite values) being given either by Δ or by the closure $\overline{\Delta}$. Moreover, we assume $F_1 \in C^{1,1}(\mathbb{R}^2)$ with

$$|\nabla F_1(s,r)| \le C(1+|s|+|r|) \quad \forall r,s \in \mathbb{R},$$
(2.3)

while $F_0 \in C^1(\Delta; [0, \infty))$, i.e., F_0 is smooth once restricted to the simplex Δ and there exists constants $c_1, c_3 > 0$ and $c_2, c_4 \ge 0$ such that

$$F_0(s,r) \ge c_1(|s|^2 + |r|^2) - c_2 \quad \forall (s,r) \in \overline{\Delta},$$
 (2.4)

and

$$\nabla F_0(s,r) \cdot (s-S,r-R)^\top \ge c_3 |\nabla F_0(s,r)| - c_4$$
 (2.5)

for all $(s,r) \neq (S,R) \in \Delta$.

(A4) The initial conditions satisfy $\varphi_{p,0}, \varphi_{d,0} \in H^1(\Omega)$ with

$$0 \le \varphi_{p,0}, \quad 0 \le \varphi_{d,0}, \quad \varphi_{p,0} + \varphi_{d,0} \le 1 \quad a.e. \quad in \ \Omega.$$

$$(2.6)$$

Moreover, the mean values

$$Y_{i,0} := \frac{1}{|\Omega|} \int_{\Omega} \varphi_{i,0}(x) \, dx \tag{2.7}$$

for i = p, d satisfy

$$(Y_{p,0}, Y_{d,0}) \in \operatorname{int} \Delta_0.$$

$$(2.8)$$

Finally, we assume that

$$F_0(\varphi_{p,0},\varphi_{d,0}) \in L^1(\Omega).$$
(2.9)

Examples. In order to clarify the above assumptions, and particularly those regarding the potential F, we introduce one example which is particularly significant and will be considered as a model case in the sequel. Namely, we consider the multi-phase logarithmic potential

$$F_0(s,r) := s \log s + r \log r + (1-s-r) \log(1-s-r),$$

$$F_1(s,r) := \frac{\chi}{2} \big(r(1-r) + s(1-s) + (1-r-s)(r+s) \big),$$
(2.10)

for a fixed positive constant χ .

Then, the assumption (2.4) is easily fulfilled by (2.10) due to the boundedness of the simplex Δ , and the assumption (2.5) is also fulfilled as we will prove in Lemma 2.2 below. Moreover, as a consequence of (2.5) we obtain by interchanging the roles of (s, r) and (S, R) an analogous inequality

$$\nabla F_0(S, R) \cdot (S - s, R - r)^\top \ge c_3 |\nabla F_0(S, R)| - c_4.$$

Then, a short computation shows that

$$(\nabla F_0(s,r) - \nabla F_0(S,R)) \cdot (s-S,r-R)^\top = \nabla F_0(s,r) \cdot (s-S,r-R)^\top + \nabla F_0(S,R) \cdot (S-s,R-r)^\top \geq c_3 |\nabla F_0(s,r)| + c_3 |\nabla F_0(S,R)| - 2c_4 \geq c_3 |\nabla F_0(s,r) - \nabla F_0(S,R)| - 2c_4.$$

$$(2.11)$$

In particular, the inequality (2.11) together with (2.4) shows that F_0 fulfills the hypotheses of [16, Prop. 2.10]. This property would be important later when we consider the Yosida approximation of F_0 for the derivation of uniform estimates.

Let us also sketch a couple of examples of functions Σ_p, Σ_d and \underline{M} that fulfill the assumption (2.2).

• First, we consider the source terms S_p , S_d introduced in (1.7), (1.8), for positive constants $\lambda_M, \lambda_A, \lambda_L$ and a bounded positive function g such that $0 < g(s) \leq 1$. For instance, one can take $g(s) = \max(n_c, \min(s, 1))$ for some constant $n_c \in (0, 1)$. Note that with this choice one can take $K_{p,-} = 0$, $K_{p,-} = \lambda_M$, $K_{d,-} = K_{d,+} = 0$. In what follows the mean of a function f over Ω is denoted as $(f)_{\Omega}$. Then, integrating (1.6a) and (1.6b) over Ω , and applying the boundary conditions (1.9) leads to the following ODE system for the mean values $Y_p := (\varphi_p)_{\Omega}$ and $Y_d := (\varphi_d)_{\Omega}$:

$$\frac{d}{dt} \begin{pmatrix} Y_p \\ Y_d \end{pmatrix} = \begin{pmatrix} -\lambda_A & 0 \\ \lambda_A & -\lambda_L \end{pmatrix} \begin{pmatrix} Y_p \\ Y_d \end{pmatrix} + \begin{pmatrix} \lambda_M(g(n))_{\Omega} \\ 0 \end{pmatrix}.$$

The matrix \underline{M} is invertible with eigenvalues $\{-\lambda_A, -\lambda_L\}$, hence the fixed point

$$\begin{pmatrix} Y_p^* \\ Y_d^* \end{pmatrix} = -\underline{\underline{M}}^{-1} \begin{pmatrix} \lambda_M(g(n))_{\Omega} \\ 0 \end{pmatrix} = \begin{pmatrix} \frac{\lambda_M}{\lambda_A}(g(n))_{\Omega} \\ \frac{\lambda_M}{\lambda_L}(g(n))_{\Omega} \end{pmatrix}$$

is asymptotically stable. Under the following constraints on the rates:

$$\lambda_M(\lambda_A + \lambda_L) < \lambda_A \lambda_L, \quad \lambda_A < 2\lambda_L,$$

we can easily show that (Y_p^*, Y_d^*) lies in the interior of the simplex Δ , and (2.2) holds when we take Δ_0 to be a ball centered at (Y_p^*, Y_d^*) with sufficiently small radius $\eta > 0$. Indeed, thanks to $n_c > 0$ we easily see that $Y_p^*, Y_d^* > 0$, while using $g \leq 1$ shows that

$$Y_p^* + Y_d^* \le \lambda_M \left(\frac{1}{\lambda_A} + \frac{1}{\lambda_L}\right) < 1$$

when we assume the hypothesis $\lambda_M(\lambda_A + \lambda_L) < \lambda_A \lambda_L$. Furthermore, taking a parameterization of the circle $\partial \Delta_0$ as $(\eta \cos \theta + Y_p^*, \eta \sin \theta + Y_d^*)$ for $\theta \in [0, 2\pi]$ with normal $\boldsymbol{n} = (\cos \theta, \sin \theta)$, a short computation shows that

$$\begin{bmatrix} \underline{M} \begin{pmatrix} \eta \cos \theta + Y_p^* \\ \eta \sin \theta + Y_d^* \end{pmatrix} + \begin{pmatrix} \lambda_M(g(n))_{\Omega} \\ 0 \end{pmatrix} \end{bmatrix} \cdot \begin{pmatrix} \cos \theta \\ \sin \theta \end{pmatrix}$$
$$= -\lambda_A \eta \cos^2 \theta + \lambda_A \eta \cos \theta \sin \theta - \lambda_L \eta \sin^2 \theta$$
$$\leq -\frac{\lambda_A}{2} \eta \cos^2 \theta - \left(\lambda_L - \frac{\lambda_A}{2}\right) \eta \sin^2 \theta \leq -\frac{1}{2} \min(\lambda_A, 2\lambda_L - \lambda_A) \eta < 0$$

under the assumption $2\lambda_L > \lambda_A$.

As a second model case, for λ > 0 we take <u>M</u> as −λ times the identity matrix (a more general negative definite diagonal matrix could also be considered) and

$$\Sigma(\varphi_p, \varphi_d, n) = \mathbf{k} + \Sigma_0(\varphi_p, \varphi_d, n), \qquad (2.12)$$

where $\boldsymbol{k} = (\lambda/3, \lambda/3)$ and Σ_0 is a C^1 function of its arguments such that

$$|\Sigma_0||_{L^\infty(\mathbb{R}^3;\mathbb{R}^2)} \le K \tag{2.13}$$

for some K > 0. This corresponds in fact to

$$\begin{pmatrix} S_p \\ S_d \end{pmatrix} = \underline{\underline{M}} \begin{pmatrix} \varphi_p - 1/3 \\ \varphi_d - 1/3 \end{pmatrix} + \Sigma_0(\varphi_p, \varphi_d, n)$$
(2.14)

and in particular we can take $K_{p,-} = K_{d,-} = \lambda/3 - K$ and $K_{p,+} = K_{d,+} = \lambda/3 + K$. Note also that the point (1/3, 1/3) can be seen as the "center" of the simplex (indeed it represents the configuration where all the species have the same proportion). Hence, here we are decomposing (S_p, S_d) as the sum of an affine part that tends to keep the configuration close to the center of the simplex, plus the perturbation Σ_0 .

With this choice, we now check that, at least if λ is large enough (depending on K), then there exists $\varepsilon > 0$ such that

$$Y_i = \varepsilon \Rightarrow Y'_i > 0, \qquad (1 - Y_p - Y_d) = \varepsilon \Rightarrow Y'_p + Y'_d < 0.$$

Indeed, let $a \in [-K, K]$. Then, for $Y_i = \varepsilon$ we have

$$-\lambda\left(Y_i - \frac{1}{3}\right) + a = \lambda\left(\frac{1}{3} - \varepsilon\right) + a \ge \lambda\left(\frac{1}{3} - \varepsilon\right) - K$$

which, for $\varepsilon < 1/3$, is greater than 0 if λ is large enough compared to K. Analogously, for $a, b \in [-K, K]$ and $Y_p + Y_d = 1 - \varepsilon$,

$$-\lambda\left(Y_p + Y_d - \frac{2}{3}\right) + a + b = -\lambda\left(\frac{1}{3} - \varepsilon\right) + a + b \le -\lambda\left(\frac{1}{3} - \varepsilon\right) + 2K$$

which is negative under the same conditions as before. Consequently, one can take Δ_0 as the set $\{Y_p \geq \varepsilon, Y_d \geq \varepsilon, Y_p + Y_d \leq 1 - \varepsilon\}$, which is larger for ε closer to 0. Namely, Δ_0 is closer to Δ if the constant λ is big compared to the L^{∞} -norm of the perturbation Σ_0 . To be more precise, we have to remark that the above choice of Δ_0 is nonsmooth (and the normal \boldsymbol{n} is not defined in the vertices). On the other hand it is easy to check that taking a smaller Δ_0 whose vertices are smoothed out the above computations are still effective.

We can now define a suitable notion of weak solution to the initial-boundary value problem for system (1.6a)-(1.6g):

Definition 2.1. We say that a multiple $(\varphi_p, \mu_p, \eta_p, \varphi_d, \mu_d, \eta_d, \boldsymbol{u}, q, n)$ is a weak solution to the multi-species tumor model (1.6) over the interval (0,T) if

(1) the following regularity properties hold:

$$\varphi_i \in H^1(0,T; H^1(\Omega)') \cap L^{\infty}(0,T; H^1(\Omega)) \cap L^2(0,T; H^2(\Omega)), \qquad (2.15a)$$

with $0 \le \varphi_i \le 1$, $\varphi_p + \varphi_d \le 1$ a.e. in Q ,

$$\mu_i \in L^2(0, T; H^1(\Omega)),$$
(2.15b)

$$\eta_i \in L^2(Q), \tag{2.15c}$$

$$\boldsymbol{u} \in L^2(Q) \text{ with } \operatorname{div} \boldsymbol{u} \in L^2(Q),$$

$$(2.15d)$$

$$q \in L^2(0, T; H^1_0(\Omega)),$$
 (2.15e)

$$n \in (1 + L^2(0, T; H^2(\Omega) \cap H^1_0(\Omega))), \quad 0 \le n \le 1 \text{ a.e. in } Q,$$
(2.15f)

for i = p, d.

(2) Equations (1.6a)-(1.6g) hold, for a.e. $t \in (0,T)$ and for i = p, d, in the following

weak sense:

$$\langle \partial_t \varphi_i, \zeta \rangle + \int_{\Omega} M_i \nabla \mu_i \cdot \nabla \zeta - \varphi_i \boldsymbol{u} \cdot \nabla \zeta \, d\boldsymbol{x} = \int_{\Omega} S_i \zeta \, d\boldsymbol{x} \quad \forall \zeta \in H^1(\Omega), \quad (2.16a)$$

$$\int_{\Omega} \mu_i \zeta \, dx = \int_{\Omega} \nabla \varphi_i \cdot \nabla \zeta + \eta_i \zeta + F_{1,\varphi_i}(\varphi_p, \varphi_d) \zeta \, dx \quad \forall \zeta \in H^1(\Omega), \tag{2.16b}$$

$$\int_{\Omega} \boldsymbol{u} \cdot \nabla \xi \, d\boldsymbol{x} = -\int_{\Omega} (S_p + S_d) \xi \, d\boldsymbol{x} \quad \forall \xi \in H_0^1(\Omega), \tag{2.16c}$$

$$\int_{\Omega} \boldsymbol{u} \cdot \boldsymbol{\zeta} \, d\boldsymbol{x} = \int_{\Omega} -\nabla q \cdot \boldsymbol{\zeta} - \varphi_p \nabla \mu_p \cdot \boldsymbol{\zeta} - \varphi_d \nabla \mu_d \cdot \boldsymbol{\zeta} \, d\boldsymbol{x} \quad \forall \boldsymbol{\zeta} \in (L^2(\Omega))^d, \quad (2.16d)$$

$$0 = -\Delta n + \varphi_p n \quad a.e. \ in \ \Omega, \tag{2.16e}$$

$$\eta_i = F_{0,\varphi_i}(\varphi_p, \varphi_d) \quad a.e. \text{ in } \Omega, \tag{2.16f}$$

$$S_p = \Sigma_p(n, \varphi_p, \varphi_d) + m_{pp}\varphi_p + m_{pd}\varphi_d \quad a.e. \text{ in } \Omega,$$
(2.16g)

$$S_d = \Sigma_d(n, \varphi_p, \varphi_d) + m_{dp}\varphi_p + m_{dd}\varphi_d \quad a.e. \text{ in } \Omega.$$
(2.16h)

Moreover, there hold the initial conditions

$$\varphi_p(x,0) = \varphi_{p,0}(x), \quad \varphi_d(x,0) = \varphi_{d,0}(x) \quad a.e. \text{ in } \Omega, \tag{2.16i}$$

where $\langle \cdot, \cdot \rangle$ denotes the duality pairing between $H^1(\Omega)$ and its dual $H^1(\Omega)'$.

It is worth noting that now the first two boundary conditions in (1.9) have been incorporated in the weak formulations (2.16a), (2.16b). Moreover, the boundary conditions q = 0and n = 1 a.e. on Σ are built into the function spaces in (2.15e) and (2.15f). Furthermore, the attainment of the initial conditions (2.16i) is due to the continuous embedding

$$H^{1}(0,T; H^{1}(\Omega)') \cap L^{\infty}(0,T; H^{1}(\Omega)) \subset C^{0}([0,T]; L^{2}(\Omega)),$$

and thus the initial conditions (2.16i) makes sense as equalities in the space $L^2(\Omega)$. Finally, it is worth saying some words about the auxiliary variables η_p , η_d . Actually, using the language of convex analysis, relations (2.16f) for i = p, d may be equivalently stated by saying that the vector $\boldsymbol{\eta} = (\eta_p, \eta_d)$ belongs at almost every point $(x, t) \in Q$ to the subdifferential $\partial F_0(\varphi_p, \varphi_d)$ which is a maximal monotone graph in $\mathbb{R}^2 \times \mathbb{R}^2$. In principle such an object may be a multivalued mapping; here, however, in view of the fact that F_0 is assumed to be smooth in Δ (cf. (A3)), ∂F_0 may be simply identified with the gradient ∇F_0 . On the other hand, the use of some techniques from convex analysis and monotone operators will be required in the last part of the proof.

We are now ready to state the main result of this paper

Theorem 2.1. Let the hypotheses stated in Assumption 2.1 hold. Then there exists at least one weak solution $(\varphi_p, \mu_p, \eta_p, \varphi_d, \mu_d, \eta_d, \boldsymbol{u}, q, n)$ to the multi-species tumor model (1.6) in the sense of Definition 2.1.

Before we prove our main result, we show that the convex part F_0 of the model potential (2.10) satisfies the assumption (2.5).

Lemma 2.2. Let F_0 be defined as

$$F_0(s,r) = s \log s + r \log r + (1 - s - r) \log(1 - s - r)$$

and let Δ_0 be a compact subset of Δ . Then there exist positive constants c_*, C_* depending only on Δ_0 such that (2.5) holds.

Proof. For any $(S, R) \in \Delta_0$, computing the gradient of F_0 leads to

$$\nabla F_0(s,r) \cdot (s-S,r-R)^{\top} = (s-S)\log s + (r-R)\log r + ((R+S) - (r+s))\log(1 - (r+s)).$$

For $s, S \in (0, 1)$ it holds that

$$(s-S)\log s = \begin{cases} >0 & \text{if } s < S, \\ <0 & \text{if } s > S, \\ =0 & \text{if } s = S, \end{cases} \text{ and } (s-S)\log s \to \begin{cases} 0 & \text{as } s \to 1, \\ \infty & \text{as } s \to 0. \end{cases}$$

In particular, the function $(s-S)\log(s)$ is bounded from below by some negative constant. Hence, there exists a constant $d_1 \ge 0$ such that

$$(s-S)\log s \ge \frac{S}{2}|\log s| - d_1,$$

and it is clear that as $S \in (0, 1)$ we can choose d_1 independent of S. In a similar fashion, there exists a constant $d_2 \ge 0$ (that can be chosen independent of R) such that

$$(r-R)\log r \ge \frac{R}{2}|\log r| - d_2.$$

Lastly, as $(s, r) \in \Delta$ we have $r + s \in (0, 1)$ and consequently there exists a constant $d_3 \ge 0$ independent of R, S such that

$$((R+S) - (r+s))\log(1 - (r+s)) \ge \frac{1 - (R+S)}{2}|\log(1 - (r+s))| - d_3$$

Summing the above then yields

$$\nabla F_0(s,r) \cdot (s-S,r-R)^\top$$

$$\geq \frac{1}{2}\min(R,S,1-(R+S)) (|\log r|+|\log s|+|\log(1-(r+s))|) - C(d_1,d_2,d_3)$$

$$\geq \frac{1}{4}\min(R,S,1-(R+S)) |\nabla F_0(s,r)| - C(d_1,d_2,d_3).$$

Now for $(R, S) \in \Delta_0$, we see that

$$\min(R, S, 1 - (R + S)) \ge c_* > 0$$

for some constant $c_* > 0$ depending only on Δ_0 . This concludes the proof of Lemma 2.2.

3 Approximation scheme

In order to start our existence proof, we introduce a regularized version of our model. First of all, for $\varepsilon \in (0, 1)$ intended to go to 0 in the limit, we consider a convex function

$$F_{\varepsilon} : \mathbb{R}^2 \to [0, +\infty) \tag{3.1}$$

such that, for any $\varepsilon \in (0,1)$, ∇F_{ε} is globally Lipschitz continuous. Moreover, we assume that F_{ε} converges in a suitable sense to F_0 as $\varepsilon \searrow 0$. Various choices are possible for F_{ε} ,

but in light of the analysis below, we take F_{ε} as the Moreau–Yosida approximation of F_0 (cf. [6]), which is defined as

$$F_{\varepsilon}(s,r) := \min_{(p,q) \in \mathbb{R}^2} \left(\frac{1}{2\varepsilon} |(p-s,q-r)|^2 + F_0(p,q) \right) \quad \text{for } \varepsilon \in (0,1)$$

It is well-known that F_{ε} is convex and differentiable with derivative ∇F_{ε} that is globally Lipschitz continuous with Lipschitz constant scaling with $\frac{1}{\varepsilon}$. More importantly, thanks to the fact that F_0 satisfies (2.11), it turns out that F_0 fulfills the hypothesis of [16, Prop. 2.10], whence, by [16, Prop. 2.13], there exist positive constants c_*, C_* such that

$$c_* |\nabla F_{\varepsilon}(s, r) - \nabla F_{\varepsilon}(S, R)| \le (\nabla F_{\varepsilon}(s, r) - \nabla F_{\varepsilon}(S, R)) \cdot (s - S, r - R)^{\top} + C_*$$
(3.2)

for all $(s,r) \neq (S,R) \in \mathbb{R}^2$. In particular an analogue of (2.11) also holds for F_{ε} with constants c_*, C_* independent of $\varepsilon \in (0, 1)$.

A difficulty concerned with the regularization of F_0 stands in the fact that F_{ε} is no longer a singular function; consequently, the uniform boundedness properties $0 \leq \varphi_p$, $0 \leq \varphi_d$, $\varphi_p + \varphi_d \leq 1$ are not expected to hold in the approximation. For this reason, in order that the a priori estimates still work, some terms have to be truncated in the regularized system. In addition to that, we also include a number of regularizing terms depending by a further parameter $\delta > 0$ which is intended to go to 0 in the limit. Finally, we remove the explicit dependence on \boldsymbol{u} in the equations and rewrite the transport terms in (1.6a), (1.6b) directly in terms of the pressure q. Hence, introducing the cutoff operator

$$T(r) := \max\{0, \min\{1, r\}\},\$$

our regularized system takes the form

$$\partial_t \varphi_p = M_p \Delta \mu_p + \operatorname{div} \left(T(\varphi_p) \nabla q \right) + \operatorname{div} \left(T(\varphi_p)^2 \nabla \mu_p + T(\varphi_p) T(\varphi_d) \nabla \mu_d \right) + S_p, \quad (3.3a)$$

$$\mu_p = -\delta \Delta \partial_t \varphi_p + F_{\varepsilon,p}(\varphi_p, \varphi_d) + F_{1,p}(\varphi_p, \varphi_d) - \Delta \varphi_p, \tag{3.3b}$$

$$\partial_t \varphi_d = M_d \Delta \mu_d + \operatorname{div} \left(T(\varphi_d) \nabla q \right) + \operatorname{div} \left(T(\varphi_p) T(\varphi_d) \nabla \mu_p + T(\varphi_d)^2 \nabla \mu_d \right) + S_d, \quad (3.3c)$$

$$\mu_d = -\delta \Delta \partial_t \varphi_d + F_{del} \left(\varphi_d - \varphi_d \right) + F_{del} \left(\varphi_d - \varphi_d \right) + S_d, \quad (3.3c)$$

$$\mu_d = -\delta \Delta \partial_t \varphi_d + F_{\varepsilon,d}(\varphi_p, \varphi_d) + F_{1,d}(\varphi_p, \varphi_d) - \Delta \varphi_d,$$

$$S_n = \sum_p (n, \varphi_n, \varphi_d) + m_{np} \varphi_n + m_{nd} \varphi_d.$$
(3.3e)

$$S_p = \Sigma_p(n, \varphi_p, \varphi_d) + m_{pp}\varphi_p + m_{pd}\varphi_d,$$

$$S_d = \Sigma_d(n, \varphi_p, \varphi_d) + m_{dn}\varphi_n + m_{dd}\varphi_d,$$
(3.3f)

$$\delta\partial_t q = \Delta q - \delta\Delta^2 q + \operatorname{div}\left(T(\varphi_n)\nabla\mu_n + T(\varphi_d)\nabla\mu_d\right) + S_n + S_d, \tag{3.3g}$$

$$0 = -\Delta n + T(\varphi_p)n, \tag{3.3h}$$

furnished with the initial and boundary conditions

$$\varphi_p(0) = \varphi_{p,0,\delta}, \quad \varphi_d(0) = \varphi_{d,0,\delta}, \quad q(0) = 0 \text{ in } \Omega, \tag{3.4a}$$

$$M_i \partial_{\boldsymbol{n}} \mu_i + T(\varphi_i) (\nabla q + T(\varphi_p) \nabla \mu_p + T(\varphi_d) \nabla \mu_d) \cdot \boldsymbol{n} = 0 \text{ on } \Gamma, \qquad (3.4b)$$

$$\partial_{\boldsymbol{n}}\varphi_i = 0, \quad \boldsymbol{n} = 1, \quad \boldsymbol{q} = 0, \quad \Delta \boldsymbol{q} = 0 \text{ on } \boldsymbol{\Gamma}, \quad (3.4c)$$

where for each $\delta > 0$, i = p, d, the initial data $\varphi_{i,0,\delta} \in H^2_n(\Omega)$ is defined as the solution f_i to

$$-\delta\Delta f_i + f_i = \varphi_{i,0} \text{ in } \Omega, \quad \partial_{\boldsymbol{n}} f_i = 0 \text{ on } \Gamma.$$
(3.5)

We have used here the notation $H^2_n(\Omega)$ for the space of $H^2(\Omega)$ -functions satisfying homogeneous Neumann boundary condition on Γ . Then, it is well-known that, for each $\delta \in (0,1], f_i \in H_n^2(\Omega)$. More precisely, testing (3.5) by f_i and $-\Delta f_i$, respectively, one obtains

$$2\delta \|\nabla f_i\|_{L^2(\Omega)}^2 + \|f_i\|_{L^2(\Omega)}^2 \le \|\varphi_{i,0}\|_{L^2(\Omega)}^2,$$

$$2\delta \|\Delta f_i\|_{L^2(\Omega)}^2 + \|\nabla f_i\|_{L^2(\Omega)}^2 \le \|\nabla \varphi_{i,0}\|_{L^2(\Omega)}^2.$$

(3.6)

Furthermore, elliptic regularity arguments yield the additional estimate

$$\|f_i\|_{H^2(\Omega)} \le C\left(\|\Delta f_i\|_{L^2(\Omega)} + \|f\|_{L^2(\Omega)}\right) \le C\left(1 + \delta^{-\frac{1}{2}}\right)\|\varphi_{i,0}\|_{H^1(\Omega)}.$$
(3.7)

3.1 Auxiliary Cahn–Hilliard problem

Fix now $\bar{q} \in L^2(0,T; H^1(\Omega))$ and $\bar{n} \in L^2(Q)$ with $0 \leq \bar{n} \leq 1$ almost everywhere in Q. Then, we first consider

$$\partial_t \varphi_p = M_p \Delta \mu_p + \operatorname{div} \left(T(\varphi_p) \nabla \bar{q} \right) + \operatorname{div} \left(T(\varphi_p)^2 \nabla \mu_p + T(\varphi_p) T(\varphi_d) \nabla \mu_d \right) + S_p, \quad (3.8a)$$

$$\mu_p = -\delta \Delta \partial_t \varphi_p + F_{\varepsilon,p}(\varphi_p, \varphi_d) + F_{1,p}(\varphi_p, \varphi_d) - \Delta \varphi_p, \qquad (3.8b)$$

$$\partial_t \varphi_d = M_d \Delta \mu_d + \operatorname{div} \left(T(\varphi_d) \nabla \bar{q} \right) + \operatorname{div} \left(T(\varphi_p) T(\varphi_d) \nabla \mu_p + T(\varphi_d)^2 \nabla \mu_d \right) + S_d, \quad (3.8c)$$

$$\mu_d = -\delta \Delta \partial_t \varphi_d + F_{\varepsilon,d}(\varphi_p, \varphi_d) + F_{1,d}(\varphi_p, \varphi_d) - \Delta \varphi_d, \qquad (3.8d)$$

$$S_p = \Sigma_p(\bar{n}, \varphi_p, \varphi_d) + m_{pp}\varphi_p + m_{pd}\varphi_d, \qquad (3.8e)$$

$$S_d = \Sigma_d(\bar{n}, \varphi_p, \varphi_d) + m_{dp}\varphi_p + m_{dd}\varphi_d, \qquad (3.8f)$$

complemented with the initial and boundary conditions (3.4a)-(3.4c). The above is a Cahn–Hilliard system with source term. Note that \bar{q} and \bar{n} are given. Existence of a solution can be proved for instance via a Galerkin approximation, and we will only derive the necessary a priori estimates.

Lemma 3.1. For each $\varepsilon \in (0,1)$, $\delta \in (0,1)$, suppose (2.1) holds, and $F_{\varepsilon} : \mathbb{R}^2 \to [0,+\infty)$ and $F_1 : \mathbb{R}^2 \to \mathbb{R}$ are given such that ∇F_{ε} , ∇F_1 are globally Lipschitz continuous. Then, for given $\bar{q} \in L^2(0,T; H^1(\Omega))$ and $\bar{n} \in L^2(Q)$, there exists a unique weak solution $(\varphi_p, \mu_p, \varphi_d, \mu_d)$ to (3.8) in the following sense:

(1) the functions have the following regularity properties:

$$\varphi_i \in H^1(0, T; H^2(\Omega)), \quad \mu_i \in L^2(0, T; H^1(\Omega)),$$

with

$$\varphi_i(0) = \varphi_{0,i,\delta} \text{ in } \Omega.$$

(2) Equations (3.8b), (3.8d), (3.8e) and (3.8f) hold a.e. in Q, and equations (3.8a) and (3.8c) hold for a.e. $t \in (0,T)$ in the following weak sense:

$$0 = \int_{\Omega} (\partial_t \varphi_p - S_p) \zeta + (M_p \nabla \mu_p + T(\varphi_p) \nabla \bar{q} + T(\varphi_p)^2 \nabla \mu_p + T(\varphi_p) T(\varphi_d) \nabla \mu_d) \cdot \nabla \zeta \, dx,$$

$$0 = \int_{\Omega} (\partial_t \varphi_d - S_d) \zeta + (M_d \nabla \mu_d + T(\varphi_d) \nabla \bar{q} + T(\varphi_d)^2 \nabla \mu_d + T(\varphi_p) T(\varphi_d) \nabla \mu_p) \cdot \nabla \zeta \, dx,$$

for all $\zeta \in H^1(\Omega).$

Proof. In the following, the symbol C denotes positive constants that depend only on the given parameters of the problem and in particular are independent of φ_p , μ_p , φ_d , μ_d .

First estimate. Testing (3.8a) with μ_p , (3.8b) with $\partial_t \varphi_p$, (3.8c) with μ_d and (3.8d) with $\partial_t \varphi_d$, and summing yields

$$\frac{d}{dt}E_{\varepsilon}(\varphi_{p},\varphi_{d}) + M_{p}\|\nabla\mu_{p}\|_{L^{2}(\Omega)}^{2} + M_{d}\|\nabla\mu_{d}\|_{L^{2}(\Omega)}^{2}
+ \delta\|\nabla\partial_{t}\varphi_{p}\|_{L^{2}(\Omega)}^{2} + \delta\|\nabla\partial_{t}\varphi_{d}\|_{L^{2}(\Omega)}^{2} + \int_{\Omega}|T(\varphi_{p})\nabla\mu_{p} + T(\varphi_{d})\nabla\mu_{d}|^{2} dx \qquad (3.9)$$

$$= \int_{\Omega}(S_{p}\mu_{p} + S_{d}\mu_{d}) dx - \int_{\Omega}\left(T(\varphi_{p})\nabla\bar{q}\cdot\nabla\mu_{p} + T(\varphi_{d})\nabla\bar{q}\cdot\nabla\mu_{d}\right) dx.$$

Note that the additional (nonnegative) contribution $\int_{\Omega} |T(\varphi_d)\nabla\mu_p + T(\varphi_d)\nabla\mu_d|^2 dx$ comes from the fact that the equations have been restated in terms of the pressure q. Note also that, here, in view of the fact that the potential is regularized, the approximate energy E_{ε} is given by

$$E_{\varepsilon} = \frac{1}{2} \left(\left\| \nabla \varphi_p \right\|_{L^2(\Omega)}^2 + \left\| \nabla \varphi_d \right\|_{L^2(\Omega)}^2 \right) + \int_{\Omega} \left(F_{\varepsilon}(\varphi_p, \varphi_d) + F_1(\varphi_p, \varphi_d) \right) dx$$

and it may be no longer coercive with respect to φ_p, φ_d . For this reason, we need to add to (3.9) the product of (3.8a) by φ_p and the product of (3.8c) by φ_d . By standard computations the sum of these contributions can be written in the form of the following differential inequality:

$$\frac{1}{2} \frac{d}{dt} \left(\|\varphi_p\|_{L^2(\Omega)}^2 + \|\varphi_d\|_{L^2(\Omega)}^2 \right) \\
\leq \sigma \left(\|\nabla \mu_p\|_{L^2(\Omega)}^2 + \|\nabla \mu_d\|_{L^2(\Omega)}^2 \right) + c_\sigma \left(\|\nabla \varphi_p\|_{L^2(\Omega)}^2 + \|\nabla \varphi_d\|_{L^2(\Omega)}^2 \right) \\
+ C \left(1 + \|\nabla \bar{q}\|_{L^2(\Omega)}^2 + \|\varphi_p\|_{L^2(\Omega)}^2 + \|\varphi_d\|_{L^2(\Omega)}^2 \right)$$
(3.10)

for small constant $\sigma > 0$ and correspondingly large constant c_{σ} . Hence, the sum of (3.9) and large K > 0 times (3.10) gives

$$\frac{d}{dt} \Big(E_{\varepsilon}(\varphi_{p},\varphi_{d}) + \frac{K}{2} \Big(\|\varphi_{p}\|_{L^{2}(\Omega)}^{2} + \|\varphi_{d}\|_{L^{2}(\Omega)}^{2} \Big) \Big)
+ (M_{p} - K\sigma) \|\nabla\mu_{p}\|_{L^{2}(\Omega)}^{2} + (M_{d} - K\sigma) \|\nabla\mu_{d}\|_{L^{2}(\Omega)}^{2}
+ \delta \|\nabla\partial_{t}\varphi_{p}\|_{L^{2}(\Omega)}^{2} + \delta \|\nabla\partial_{t}\varphi_{d}\|_{L^{2}(\Omega)}^{2}
\leq \int_{\Omega} (S_{p}\mu_{p} + S_{d}\mu_{d}) dx - (T(\varphi_{p})\nabla\bar{q} \cdot \nabla\mu_{p} + T(\varphi_{d})\nabla\bar{q} \cdot \nabla\mu_{d}) dx
+ Kc_{\sigma} \Big(\|\nabla\varphi_{p}\|_{L^{2}(\Omega)}^{2} + \|\nabla\varphi_{d}\|_{L^{2}(\Omega)}^{2} \Big)
+ KC \Big(1 + \|\nabla\bar{q}\|_{L^{2}(\Omega)}^{2} + \|\varphi_{p}\|_{L^{2}(\Omega)}^{2} + \|\varphi_{d}\|_{L^{2}(\Omega)}^{2} \Big).$$
(3.11)

Now, we first take K large enough so that the modified energy is coercive, namely

$$E_{\varepsilon} + \frac{K}{2} \left(\|\varphi_p\|_{L^2(\Omega)}^2 + \|\varphi_d\|_{L^2(\Omega)}^2 \right) \ge k \left(\|\varphi_p\|_{H^1(\Omega)}^2 + \|\varphi_d\|_{H^1(\Omega)}^2 \right) - C, \tag{3.12}$$

for some k > 0 (note that k can be chosen independently of ε ; indeed, by (3.1), F_{ε} is non-negative, and the smooth non-convex part F_1 has at most quadratic growth). Then, after K is fixed, we also take σ so small that $\min\{M_p - K\sigma, M_d - K\sigma\} \ge \kappa$ for some constant $\kappa > 0$. It remains to control the integral terms on the right-hand side of (3.11). Observe that

$$\left| \int_{\Omega} \left(T(\varphi_p) \nabla \bar{q} \cdot \nabla \mu_p + T(\varphi_d) \nabla \bar{q} \cdot \nabla \mu_d \right) dx \right|$$

$$\leq \frac{\kappa}{4} \left(\|\nabla \mu_p\|_{L^2(\Omega)}^2 + \|\nabla \mu_d\|_{L^2(\Omega)}^2 \right) + C \|\nabla \bar{q}\|_{L^2(\Omega)}^2,$$

and recalling $(f)_{\Omega}$ denotes the mean value of f over Ω , we have

$$\int_{\Omega} \left(S_{p}(\mu_{p} - (\mu_{p})_{\Omega}) + S_{p}(\mu_{p})_{\Omega} \right) dx
= \int_{\Omega} S_{p}(\mu_{p} - (\mu_{p})_{\Omega}) dx + (\mu_{p})_{\Omega} \int_{\Omega} S_{p} dx
\leq \frac{\kappa}{4} \|\nabla \mu_{p}\|_{L^{2}(\Omega)}^{2} + C \left(1 + \|\varphi_{p}\|_{L^{2}(\Omega)}^{2} + \|\varphi_{d}\|_{L^{2}(\Omega)}^{2} \right)
+ C_{\varepsilon} \left(1 + \|\varphi_{p}\|_{L^{2}(\Omega)}^{2} + \|\varphi_{d}\|_{L^{2}(\Omega)}^{2} \right),$$
(3.13)

in view of the Lipschitz regularity of $F_{\varepsilon,p}$ and $F_{1,p}$. The term $S_d \mu_d$ is controlled analogously. Then, collecting the above computations, (3.11) becomes

$$\frac{d}{dt} \left(E_{\varepsilon}(\varphi_{p},\varphi_{d}) + \frac{K}{2} \left(\|\varphi_{p}\|_{L^{2}(\Omega)}^{2} + \|\varphi_{d}\|_{L^{2}(\Omega)}^{2} \right) \right)
+ \frac{\kappa}{2} \left(\|\nabla\mu_{p}\|_{L^{2}(\Omega)}^{2} + \|\nabla\mu_{d}\|_{L^{2}(\Omega)}^{2} \right) + \delta \|\nabla\partial_{t}\varphi_{p}\|_{L^{2}(\Omega)}^{2} + \delta \|\nabla\partial_{t}\varphi_{d}\|_{L^{2}(\Omega)}^{2}
\leq C_{\varepsilon} \left(1 + \|\varphi_{p}\|_{H^{1}(\Omega)}^{2} + \|\varphi_{d}\|_{H^{1}(\Omega)}^{2} \right) + C \|\nabla\bar{q}\|_{L^{2}(\Omega)}^{2},$$
(3.14)

for some positive constants C, C_{ε} that are independent of δ . By virtue of the coercivity property (3.12) and a Gronwall argument then yields

$$\|\varphi_p\|_{L^{\infty}(0,T;H^1(\Omega))} + \|\varphi_d\|_{L^{\infty}(0,T;H^1(\Omega))} \le C_{\varepsilon},$$
(3.15)

$$\|\mu_p\|_{L^2(0,T;H^1(\Omega))} + \|\mu_d\|_{L^2(0,T;H^1(\Omega))} \le C_{\varepsilon}, \tag{3.16}$$

$$\|\nabla \partial_t \varphi_p\|_{L^2(0,T;L^2(\Omega))} + \|\nabla \partial_t \varphi_d\|_{L^2(0,T;L^2(\Omega))} \le C_{\varepsilon,\delta}.$$
(3.17)

Second estimate. Testing (3.8b) with $-\Delta \partial_t \varphi_p$ and (3.8d) with $-\Delta \partial_t \varphi_d$ and using that $\Delta \varphi_{i,0,\delta} \in L^2(\Omega)$, we obtain

$$\|\Delta\varphi_p\|_{H^1(0,T;L^2(\Omega))} + \|\Delta\varphi_d\|_{H^1(0,T;L^2(\Omega))} \le C_{\varepsilon,\delta},$$

and by elliptic regularity we get the following additional estimate:

$$\|\varphi_p\|_{H^1(0,T;H^2(\Omega))} + \|\varphi_d\|_{H^1(0,T;H^2(\Omega))} \le C_{\varepsilon,\delta}.$$
(3.18)

Uniqueness. Let us denote by $\hat{\varphi}_p$, $\hat{\varphi}_d$, $\hat{\mu}_p$ and $\hat{\mu}_d$ as the differences $\varphi_{p,1} - \varphi_{p,2}$, $\varphi_{d,1} - \varphi_{d,2}$, $\mu_{p,1} - \mu_{p,2}$ and $\mu_{d,1} - \mu_{d,2}$, respectively. Then, upon testing the difference of the equations (3.8a) by $\hat{\mu}_p$ and the difference of the equations (3.8b) by $\partial_t \hat{\varphi}_p - \Delta \hat{\varphi}_p$ leads to

$$\frac{d}{dt} \frac{1}{2} \left(\|\nabla \hat{\varphi}_p\|_{L^2(\Omega)}^2 + \delta \|\Delta \hat{\varphi}_p\|_{L^2(\Omega)}^2 \right)
+ \|\Delta \hat{\varphi}_p\|_{L^2(\Omega)}^2 + \delta \|\nabla \partial_t \hat{\varphi}_p\|_{L^2(\Omega)}^2 + \int_{\Omega} \left(M_p + (T_{p,2})^2 \right) |\nabla \hat{\mu}_p|^2 dx
= \int_{\Omega} \hat{S}_p \hat{\mu}_p - \hat{\mu}_p \Delta \hat{\varphi}_p - \hat{T}_p \nabla \bar{q} \cdot \nabla \hat{\mu}_p - \left(\hat{F}_{\varepsilon,p} + \hat{F}_{1,p} \right) \left(\partial_t \hat{\varphi}_p - \Delta \hat{\varphi}_p \right) dx
- \int_{\Omega} \nabla \hat{\mu}_p \cdot \left(\widehat{(T_p)^2} \nabla \mu_{p,1} + \left(\hat{T}_p T_{d,1} + T_{p,2} \hat{T}_d \right) \nabla \mu_{d,1} + T_{p,2} T_{d,2} \nabla \hat{\mu}_d \right) dx
=: J_1 + J_2,$$
(3.19)

where we used the notation $T_{p,1} = T(\varphi_{p,1}), \hat{T}_p = T_{p,1} - T_{p,2}, \hat{F}_{\varepsilon,p} = F_{\varepsilon,p}(\varphi_{p,1},\varphi_{d,1}) - F_{\varepsilon,p}(\varphi_{p,2},\varphi_{d,2}), \hat{S}_p = \Sigma_p(\bar{n},\varphi_{p,1},\varphi_{d,1}) - \Sigma_p(\bar{n},\varphi_{p,2},\varphi_{d,2}) + m_{pp}\hat{\varphi}_p + m_{pd}\hat{\varphi}_d, \text{ and } (T_p)^2 = \Gamma_{\varepsilon,p}(\varphi_{p,2},\varphi_{d,2}) + \Gamma_{\varepsilon,p$

 $(T_{p,1})^2 - (T_{p,2})^2 = \hat{T}_p(T_{p,1} + T_{p,2})$. Using the Lipschitz continuity of $F_{\varepsilon,p}$, $F_{1,p}$, $T(\cdot)$, Σ_i and the boundedness of $T(\cdot)$ and Σ_i , we deduce

$$\begin{split} J_{1} &\leq C \big(\|\hat{\varphi}_{p}\|_{L^{2}(\Omega)} + \|\hat{\varphi}_{d}\|_{L^{2}(\Omega)} + \|\Delta\hat{\varphi}_{p}\|_{L^{2}(\Omega)} \big) \big(\|\hat{\mu}_{p} - (\hat{\mu}_{p})_{\Omega}\|_{L^{2}(\Omega)} + |(\hat{\mu}_{p})_{\Omega}| \big) \\ &+ C \big(\|\hat{\varphi}_{p}\|_{L^{2}(\Omega)} + \|\hat{\varphi}_{d}\|_{L^{2}(\Omega)} \big) \big(\|\Delta\hat{\varphi}_{p}\|_{L^{2}(\Omega)} + \|\partial_{t}\hat{\varphi}_{p} - (\partial_{t}\hat{\varphi}_{p})_{\Omega}\|_{L^{2}(\Omega)} + |(\partial_{t}\hat{\varphi}_{p})_{\Omega}| \big), \\ &+ C \|\hat{\varphi}_{p}\|_{L^{\infty}(\Omega)} \|\nabla\hat{\mu}_{p}\|_{L^{2}(\Omega)} \|\nabla\bar{q}\|_{L^{2}(\Omega)}, \\ J_{2} &\leq C \|\nabla\hat{\mu}_{p}\|_{L^{2}(\Omega)} \big(\|\hat{\varphi}_{p}\|_{L^{\infty}(\Omega)} + \|\hat{\varphi}_{d}\|_{L^{\infty}(\Omega)} \big) \big(\|\nabla\mu_{p,1}\|_{L^{2}(\Omega)} + \|\nabla\mu_{d,1}\|_{L^{2}(\Omega)} \big) \\ &+ \frac{1}{2} \int_{\Omega} (T_{p,2})^{2} |\nabla\hat{\mu}_{p}|^{2} + (T_{d,2})^{2} |\nabla\hat{\mu}_{d}|^{2} dx. \end{split}$$

Note that by the Lipschitz property of Σ_i , $F_{\varepsilon,p}$ and $F_{1,p}$,

$$\begin{aligned} |(\partial_t \hat{\varphi}_p)_{\Omega}| &= |(\hat{S}_p)_{\Omega}| \le C \big(\|\hat{\varphi}_p\|_{L^2(\Omega)} + \|\hat{\varphi}_d\|_{L^2(\Omega)} \big), \\ |(\hat{\mu})_{\Omega}| &= |(\hat{F}_{\varepsilon,p} + \hat{F}_{1,p})_{\Omega}| \le C \big(\|\hat{\varphi}_p\|_{L^2(\Omega)} + \|\hat{\varphi}_d\|_{L^2(\Omega)} \big) \end{aligned}$$

and so, upon adding (3.19) to the corresponding equation for $\hat{\varphi}_d$ leads to

$$\frac{d}{dt} \frac{1}{2} \sum_{i=p,d} \left(\|\nabla \hat{\varphi}_i\|_{L^2(\Omega)}^2 + \delta \|\Delta \hat{\varphi}_i\|_{L^2(\Omega)}^2 \right)
+ \frac{1}{2} \sum_{i=p,d} \left(\|\Delta \hat{\varphi}_i\|_{L^2(\Omega)}^2 + \delta \|\nabla \partial_t \hat{\varphi}_i\|_{L^2(\Omega)}^2 + M_i \|\nabla \hat{\mu}_i\|_{L^2(\Omega)}^2 \right)
\leq C \left[1 + \|\nabla \bar{q}\|_{L^2(\Omega)}^2 + \sum_{\substack{i=p,d \\ j=1,2}} \|\nabla \mu_{i,j}\|_{L^2(\Omega)}^2 \right] \sum_{i=p,d} \left(\|\hat{\varphi}_i\|_{L^2(\Omega)}^2 + \|\Delta \hat{\varphi}_i\|_{L^2(\Omega)}^2 \right),$$
(3.20)

where in the above we have used the elliptic estimate

$$\|f\|_{L^{\infty}(\Omega)} \le C \|f\|_{H^{2}(\Omega)} \le C \left(\|\Delta f\|_{L^{2}(\Omega)} + \|f\|_{L^{2}(\Omega)} \right), \tag{3.21}$$

for $\hat{\varphi}_p$ and $\hat{\varphi}_d$ as they satisfy no-flux boundary conditions.

Next, we test the difference of the equations (3.8a) with $\hat{\varphi}_p$ which yields

$$\begin{split} \frac{d}{dt} \frac{1}{2} \|\hat{\varphi}_p\|_{L^2(\Omega)}^2 &= -\int_{\Omega} \nabla \hat{\varphi}_p \cdot \left(M_p \nabla \hat{\mu}_p + \hat{T}_p \nabla \bar{q} + \widehat{(T_p)^2} \nabla \mu_{p,1} + (T_{p,2})^2 \nabla \hat{\mu}_p \right) dx \\ &- \int_{\Omega} \nabla \hat{\varphi}_p \cdot \left(\left(\hat{T}_p T_{d,1} + T_{p,2} \hat{T}_d \right) \nabla \mu_{d,1} + T_{p,2} T_{d,2} \nabla \hat{\mu}_d \right) - \hat{S}_p \hat{\varphi}_p \, dx \\ &\leq C \Big(1 + \|\nabla \bar{q}\|_{L^2(\Omega)}^2 + \sum_{j=1,2} \|\nabla \mu_{p,j}\|_{L^2(\Omega)}^2 \Big) \Big(\|\nabla \hat{\varphi}_p\|_{L^2(\Omega)}^2 + \sum_{i=p,d} \|\hat{\varphi}_i\|_{L^\infty(\Omega)}^2 \Big) \\ &+ \frac{M_p}{4} \|\nabla \hat{\mu}_p\|_{L^2(\Omega)}^2 + \frac{M_d}{4} \|\nabla \hat{\mu}_d\|_{L^2(\Omega)}^2, \end{split}$$

and upon adding the analogous estimate obtained from testing (3.8c) with $\hat{\varphi}_d$ and then adding to (3.20), after applying the elliptic estimate (3.21), we arrive at the following differential inequality

$$\frac{d}{dt} \frac{1}{2} \sum_{i=p,d} \left(\|\hat{\varphi}_i\|_{L^2(\Omega)}^2 + \|\nabla\hat{\varphi}_i\|_{L^2(\Omega)}^2 + \delta \|\Delta\hat{\varphi}_i\|_{L^2(\Omega)}^2 \right) \\
+ \frac{1}{2} \sum_{i=p,d} \left(\|\Delta\hat{\varphi}_i\|_{L^2(\Omega)}^2 + \delta \|\nabla\partial_t\hat{\varphi}_i\|_{L^2(\Omega)}^2 + \frac{1}{2}M_i\|\nabla\hat{\mu}_i\|_{L^2(\Omega)}^2 \right) \\
\leq C \left[1 + \|\nabla\bar{q}\|_{L^2(\Omega)}^2 + \sum_{\substack{i=p,d\\j=1,2}} \|\nabla\mu_{i,j}\|_{L^2(\Omega)}^2 \right] \sum_{i=p,d} \left(\|\hat{\varphi}_i\|_{H^1(\Omega)}^2 + \|\Delta\hat{\varphi}_i\|_{L^2(\Omega)}^2 \right),$$

and a Gronwall argument easily entails uniqueness.

3.2 Auxiliary pressure and nutrient equations

We now consider, for $(\varphi_p, \mu_p, \varphi_d, \mu_d)$ obtained from Lemma 3.1, the following system:

$$\delta(\partial_t q + \Delta^2 q) - \Delta q = \operatorname{div} \left(T(\varphi_p) \nabla \mu_p + T(\varphi_d) \nabla \mu_d \right) + (S_p + S_d)(n, \varphi_p, \varphi_d), \quad (3.22a)$$
$$0 = -\Delta n + T(\varphi_p)n, \quad (3.22b)$$

furnished with the initial-boundary conditions resulting from (3.4a)-(3.4c).

Lemma 3.2. Let $(\varphi_p, \mu_p, \varphi_d, \mu_d)$ denote a weak solution obtained from Lemma 3.1. Then, there exists a unique pair (q, n) of solutions to (3.22) in the following sense:

(1) the functions have the following regularity properties:

$$q \in L^{2}(0,T; H^{3}(\Omega)) \cap L^{\infty}(0,T; H^{1}_{0}(\Omega)) \cap H^{1}(0,T; H^{-1}(\Omega)),$$

$$n \in L^{\infty}(0,T; W^{2,r}(\Omega)) \text{ for any } r < \infty \text{ and } 0 \le n \le 1 \text{ a.e. in } Q,$$

with

$$q(0) = 0$$
 in Ω , $q = \Delta q = 0$, $n = 1$ on Γ .

(2) Equation (3.22b) holds a.e. in Q and equation (3.22a) holds for a.e. $t \in (0,T)$ in the following weak sense:

$$0 = \delta \langle \partial_t q, \zeta \rangle + \int_{\Omega} \left(\nabla q - \delta \nabla \Delta q + T(\varphi_p) \nabla \mu_p + T(\varphi_d) \nabla \mu_d \right) \cdot \nabla \zeta \, dx$$
$$- \int_{\Omega} (S_p + S_d)(n, \varphi_p, \varphi_d) \zeta \, dx,$$

for all $\zeta \in H_0^1(\Omega)$.

Proof. We investigate the nutrient and pressure equations separately.

Nutrient equation. Since $T(\cdot)$ is bounded and non-negative, we may first consider a parabolic regularization to (3.22b), namely, we add $\gamma \partial_t n$ on the right-hand side (for $\gamma \in (0,1)$) and we complement the resulting parabolic equation (for example) with the initial condition $n_{\gamma}(0) := 1$ (which is consistent with the boundary datum). Then, applying the standard parabolic theory and the weak comparison principle it is easy to show that there exists a unique function $n_{\gamma} \in L^{\infty}(0,T;H^1(\Omega)) \cap L^2(0,T;H^2(\Omega)) \cap H^1(0,T;L^2(\Omega))$ with $0 \le n_{\gamma} \le 1$ a.e. in Q. It turns out that n_{γ} is uniformly bounded in $L^2(0,T;H^1(\Omega))$ and in passing to the limit $\gamma \to 0$ we deduce the existence of a weak solution $n \in L^2(0,T;H^1(\Omega))$ to (3.22b) with $0 \le n \le 1$ a.e. in Q. Then, as $T(\varphi_p)n \in L^{\infty}(0,T;L^{\infty}(\Omega))$, applying elliptic regularity we infer $n \in L^{\infty}(0,T;W^{2,r}(\Omega))$ for any $r < \infty$.

Pressure equation. Given n, φ_p , μ_p , φ_d and μ_d , we test (3.22a) with $q - \Delta q$. Using the boundary conditions $q = \Delta q = 0$ on Γ , we then obtain

$$\frac{d}{dt} \frac{\delta}{2} \left(\|q\|_{L^{2}(\Omega)}^{2} + \|\nabla q\|_{L^{2}(\Omega)}^{2} \right) + (1+\delta) \|\Delta q\|_{L^{2}(\Omega)}^{2} + \delta \|\nabla \Delta q\|_{L^{2}(\Omega)}^{2} + \|\nabla q\|_{L^{2}(\Omega)}^{2} \\
= \int_{\Omega} \left(T_{p} \nabla \mu_{p} + T_{d} \nabla \mu_{d} \right) \cdot \nabla \left(\Delta q - q \right) + (S_{p} + S_{d}) (q - \Delta q) \, dx \\
\leq \frac{C}{\delta} \left(\|\nabla \mu_{p}\|_{L^{2}(\Omega)}^{2} + \|\nabla \mu_{d}\|_{L^{2}(\Omega)}^{2} \right) + \frac{\delta}{2} \left(\|\nabla \Delta q\|_{L^{2}(\Omega)}^{2} + \|\nabla q\|_{L^{2}(\Omega)}^{2} \right) \\
+ \frac{C}{\delta} \left(1 + \|\varphi_{p}\|_{L^{2}(\Omega)}^{2} + \|\varphi_{d}\|_{L^{2}(\Omega)}^{2} \right) + \frac{\delta}{2} \left(\|q\|_{L^{2}(\Omega)}^{2} + \|\Delta q\|_{L^{2}(\Omega)}^{2} \right).$$
(3.23)

Integrating in time, using q(0) = 0, and applying first a Gronwall argument and then elliptic regularity leads to

$$\|q\|_{L^{\infty}(0,T;H^{1}(\Omega))} + \|q\|_{L^{2}(0,T;H^{3}(\Omega))} \le C_{\delta}.$$
(3.24)

Then, by a comparison of terms in (3.22a) we infer that

$$\|\partial_t q\|_{L^2(0,T;H^{-1}(\Omega))} \le C_\delta.$$
(3.25)

Uniqueness. Let $\hat{n} := n_1 - n_2$ and $\hat{q} := q_1 - q_2$ denote the difference between two solution pairs (q_1, n_1) and (q_2, n_2) corresponding to the same data $(\varphi_p, \mu_p, \varphi_d, \mu_d)$. Then it is straightforward to see that

$$0 = -\Delta \hat{n} + T(\varphi_p)\hat{n}, \quad \delta(\partial_t \hat{q} + \Delta^2 \hat{q}) - \Delta \hat{q} = \hat{\Sigma}_p + \hat{\Sigma}_d, \tag{3.26}$$

where for i = p, d,

$$\hat{\Sigma}_i := \Sigma_i(n_1, \varphi_p, \varphi_d) - \Sigma_i(n_2, \varphi_p, \varphi_d).$$

By testing the first equation of (3.26) with \hat{n} we easily deduce that $\hat{n} = 0$ by the Poincaré inequality. Then, testing the second equation of (3.26) with \hat{q} and noting that $\hat{\Sigma}_i = 0$ due to $n_1 = n_2$, the uniqueness of solutions is clear.

3.3 Fixed point argument

We will now apply a fixed point argument locally in time, and consider for some $T_0 \in (0, T]$ the pair $(\bar{q}, \bar{n}) \in L^2(0, T_0; H^1(\Omega)) \times L^2(0, T_0; L^2(\Omega))$ with $0 \leq \bar{n} \leq 1$ a.e. in $\Omega \times (0, T_0)$. Let us introduce the mapping $\mathcal{T} : (\bar{q}, \bar{n}) \to (q, n)$, where (q, n) is the unique solution pair to (3.22) with $(\varphi_p, \mu_p, \varphi_d, \mu_d)$ as the unique solution quadruple to (3.8). To specify the domain of \mathcal{T} we define

$$X := \left\{ (q,n) : \|q\|_{L^2(0,T_0;H^1(\Omega))} + \|n\|_{L^2(0,T_0;L^2(\Omega))} \le R, \ 0 \le n \le 1 \text{ a.e. in } \Omega \times (0,T_0) \right\},$$

where R > 0 is arbitrary but otherwise fixed. For example, one can take R = 1. Let us mention that from (3.15)-(3.16) and (3.23) one obtains the estimate

$$\|q\|_{L^{2}(0,t;H^{1}(\Omega))}^{2} \leq t \|q\|_{L^{\infty}(0,T;H^{1}(\Omega))}^{2} \leq t C_{\varepsilon,\delta,R}$$

for any $t \in (0, T]$. Similarly, since $0 \le n \le 1$ a.e. in Q, we get

$$||n||_{L^2(0,t;L^2(\Omega))}^2 \le t|\Omega|.$$

Consequently, for T_0 sufficiently small (in a way that possibly depends on ε , δ and R), we have

$$\|q\|_{L^{2}(0,T_{0};H^{1}(\Omega))} + \|n\|_{L^{2}(0,T_{0};L^{2}(\Omega))} \le C_{\varepsilon,\delta,R}T_{0}^{\frac{1}{2}} \le R.$$
(3.27)

This implies that for such a choice of T_0 , the operator \mathcal{T} maps X (which is a convex closed subset of the product Banach space $L^2(0, T_0; H^1(\Omega)) \times L^2(0, T_0; L^2(\Omega))$) into itself.

Continuity. We now aim to show that $\mathcal{T} : X \to X$ is continuous with respect to the norm of $L^2(0, T_0; H^1(\Omega)) \times L^2(0, T_0; L^2(\Omega))$, keeping in mind that thanks to the uniqueness results for the auxiliary problems (3.8) and (3.22), \mathcal{T} is a single-valued mapping. Let $(\bar{q}_k, \bar{n}_k)_{k \in \mathbb{N}} \subset X$ be a sequence that converges strongly to a limit (\bar{q}, \bar{n}) in X. We denote $(q_k, n_k) = \mathcal{T}(\bar{q}_k, \bar{n}_k)$ and $(q, n) := \mathcal{T}(\bar{q}, \bar{n})$. Then, it is easy to see that from Lemma 3.1 (more precisely (3.15)-(3.18)) there exists a corresponding sequence $(\varphi_{p,k}, \mu_{p,k}, \varphi_{d,k}, \mu_{d,k})_{k \in \mathbb{N}}$ such that

$$\|\varphi_{i,k}\|_{H^1(0,T_0;H^2(\Omega))} + \|\mu_{i,k}\|_{L^2(0,T_0;H^1(\Omega))} \le C_{\varepsilon,\delta,R}$$

for i = p, d and some constant $C = C_{\varepsilon,\delta,R}$ independent of k. Then, standard compactness results [43, §8, Cor. 4] yield

$$\varphi_{i,k} \to \varphi_i$$
 strongly in $C^0([0,T_0]; W^{1,r}(\Omega)) \cap C^0(\overline{\Omega} \times [0,T_0]),$
 $\mu_{i,k} \to \mu_i$ weakly in $L^2(0,T_0; H^1(\Omega)),$

along a non-relabelled subsequence for i = p, d, and any $r \in [1, \infty)$ in two dimensions and $r \in [1, 6)$ in three dimensions. Hence, along a non-relabelled subsequence, $\varphi_{p,k} \to \varphi_p$ uniformly in $\overline{\Omega} \times [0, T_0]$ and thus $T(\varphi_{p,k}) \to T(\varphi_p)$ uniformly in $\overline{\Omega} \times [0, T_0]$. Moreover, one can easily check that the limit functions φ_i, μ_i solve (3.8) with q, n in place of $\overline{q}, \overline{n}$. Next, taking the difference of (3.22b) for two indices a and b leads to

$$-\Delta(n_a - n_b) + (T(\varphi_{p,a}) - T(\varphi_{p,b}))n_a + T(\varphi_{p,b})(n_a - n_b) = 0,$$

and by testing with $n_a - n_b$ we obtain by the Poincaré inequality

$$\begin{aligned} \|\nabla(n_{a} - n_{b})\|_{L^{2}(0,T_{0};L^{2}(\Omega))}^{2} \\ &\leq \|n_{a} - n_{b}\|_{L^{2}(0,T_{0};L^{2}(\Omega))}\|T(\varphi_{p,a}) - T(\varphi_{p,b})\|_{L^{2}(0,T_{0};L^{2}(\Omega))} \\ &\leq C\|\nabla(n_{a} - n_{b})\|_{L^{2}(0,T_{0};L^{2}(\Omega))}\|T(\varphi_{p,a}) - T(\varphi_{p,b})\|_{L^{2}(0,T_{0};L^{2}(\Omega))} \end{aligned}$$
(3.28)

after neglecting the non-negative term $T(\varphi_{p,b})|n_a - n_b|^2$. Applying the uniform convergence of $T(\varphi_{p,k})$ we see that $\{n_k\}_{k\in\mathbb{N}}$ is a Cauchy sequence in $L^2(0, T_0; H^1(\Omega))$ and thus $n_k \to n_*$ strongly in $L^2(0, T_0; L^2(\Omega))$ for some limit function n_* . Meanwhile, from the a priori estimates (3.24)-(3.25) and standard compactness results, along a non-relabelled subsequence it holds that

$$q_k \to q_*$$
 strongly in $L^2(0,T; H^1(\Omega))$.

Let us mention here that thanks to the strong convergence of $n_k \to n_*$ in $L^2(0, T_0; L^2(\Omega))$, along a further subsequence we have a.e. convergence in $\Omega \times (0, T_0)$. Continuity of Σ_i , i = p, d, and boundedness are sufficient to ensure that the source terms $\Sigma_i(n_k, \varphi_{p,k}, \varphi_{d,k})$, i = p, d, converge to $\Sigma_i(n, \varphi_p, \varphi_d)$ strongly in $L^2(0, T_0; L^2(\Omega))$.

Hence, along a non-relabelled subsequence $\mathcal{T}(\bar{q}_k, \bar{n}_k) \to (q_*, n_*)$. On the other hand, it is easy to check that (q_*, n_*) solve (3.22) (with the limit φ_i, μ_i). Then, thanks to the uniqueness of the solutions for the auxiliary equations (3.22), one infers that, necessarily, $(q_*, n_*) = (q, n) = \mathcal{T}(\bar{q}, \bar{n})$ and the whole sequence converges. This shows the required continuity of the map \mathcal{T} .

Compactness. To apply Schauder's fixed point theorem to \mathcal{T} , it remains to show that $\mathcal{T}: X \to X$ is a compact mapping. This amounts to prove for any sequence $(\bar{q}_k, \bar{n}_k)_{k \in \mathbb{N}} \subset X$, there exists a subsequence $(\bar{q}_{k_l}, \bar{n}_{k_l})_{l \in \mathbb{N}}$ such that $(q_{k_l}, n_{k_l}) := \mathcal{T}(\bar{q}_{k_l}, \bar{n}_{k_l})$ converges

strongly to some limit (q, n) in $L^2(0, T_0; H^1(\Omega)) \times L^2(0, T_0; L^2(\Omega))$. Note that by the definition of X we have

$$||q_k||_{L^2(0,T_0;H^1(\Omega))} + ||n_k||_{L^2(0,T_0;L^2(\Omega))} \le R$$

and $0 \le n_k \le 1$ a.e. in $\Omega \times (0, T_0)$. This boundedness and a similar argument to the proof of the continuity of \mathcal{T} permit us to conclude. Indeed, by repeating the a priori estimates given above, one can easily prove that the sequence (q_k, n_k) is uniformly bounded in a better space, whence follows the desired compactness assertion.

We now state the main result of this section.

Theorem 3.3 (Local existence). Let Assumption 2.1 hold. Moreover, for each $\varepsilon \in (0, 1)$, $\delta \in (0, 1)$ let us assume that $F_{\varepsilon} : \mathbb{R}^2 \to [0, +\infty)$ and $F_1 : \mathbb{R}^2 \to \mathbb{R}$ are given such that ∇F_{ε} , ∇F_1 are globally Lipschitz continuous. Then, there exist a time $T_0 \in (0, T]$ and functions $(\varphi_p, \mu_p, \varphi_d, \mu_d, q, n)$ such that

(1) the following regularity properties

$$\begin{split} \varphi_i &\in H^1(0, T_0; H^2(\Omega)) \quad \text{for } i = p, d, \\ \mu_i &\in L^2(0, T_0; H^1(\Omega)) \quad \text{for } i = p, d, \\ q &\in L^2(0, T_0; H^3(\Omega)) \cap L^\infty(0, T_0; H_0^1(\Omega)) \cap H^1(0, T_0; H^{-1}(\Omega)), \\ n &\in L^\infty(0, T_0; W^{2, r}(\Omega)) \text{ for any } r < \infty \text{ and } 0 \le n \le 1 \text{ a.e. in } \Omega \times (0, T_0), \end{split}$$

hold together with

$$\varphi_i(0) = \varphi_{0,i,\delta}, \ i = p, d, \ q(0) = 0 \ in \ \Omega, \quad \Delta q = 0, \ n = 1 \ on \ \partial \Omega \times (0, T_0)$$

(2) Equations (3.3b), (3.3d), (3.3e), (3.3f) and (3.3h) hold a.e. in $\Omega \times (0, T_0)$, and equations (3.3a), (3.3c) and (3.3g) hold for a.e. $t \in (0, T_0)$ in the following weak sense:

$$0 = \int_{\Omega} (\partial_t \varphi_p - S_p) \zeta + (M_p \nabla \mu_p + T(\varphi_p) (\nabla q + T(\varphi_p) \nabla \mu_p + T(\varphi_d) \nabla \mu_d)) \cdot \nabla \zeta \, dx,$$

$$0 = \int_{\Omega} (\partial_t \varphi_d - S_d) \zeta + (M_d \nabla \mu_d + T(\varphi_d) (\nabla q + T(\varphi_d) \nabla \mu_d + T(\varphi_p) \nabla \mu_p)) \cdot \nabla \zeta \, dx,$$

$$0 = \delta \langle \partial_t q, \xi \rangle + \int_{\Omega} (\nabla q - \delta \nabla \Delta q + T(\varphi_p) \nabla \mu_p + T(\varphi_d) \nabla \mu_d) \cdot \nabla \xi - (S_p + S_d) \xi \, dx$$

for all $\zeta \in H^1(\Omega)$ and $\xi \in H^1_0(\Omega)$.

3.4 A priori estimates

We now derive some a priori estimates for the solution $(\varphi_p, \mu_p, \varphi_d, \mu_d, q, n)$ to (3.3) obtained from Theorem 3.3. All these estimates will be independent of the final time T_0 , which will allow us to extend the solution up to the full time interval [0, T]. For this reason, although with some abuse of notation, we shall directly work on the original time interval [0, T] and postpone the details of the extension argument to the next subsection. Below the symbol C denotes constants that are independent of δ and ε . **First estimate.** From the nutrient equation (3.3h), we obtain from the boundedness of the cut-off operator T and of n the estimate

$$\|\nabla n\|_{L^2(\Omega)}^2 + \int_{\Omega} T(\varphi_p) |n-1|^2 \, dx = -\int_{\Omega} T(\varphi_p) (n-1) \, dx \le C.$$

Hence, integrating in time yields

$$||n||_{L^2(0,T;H^1(\Omega))} \le C.$$

The weak comparison principle then yields that $0 \le n \le 1$ a.e. in $\Omega \times (0,T)$. Hence, by elliptic regularity, we arrive at

$$\|n\|_{L^{\infty}(0,T;W^{2,r}(\Omega))} \le C \quad \forall r < \infty.$$

$$(3.29)$$

Second estimate. Testing (3.3a) with μ_p , (3.3b) with $\partial_t \varphi_p$, (3.3c) with μ_d and (3.3d) with $\partial_t \varphi_d$, and summing leads to an analogous identity to (3.9) but with \bar{q} replaced by q. Then, adding the resulting identity to that obtained from testing (3.3g) with q leads to the equality

$$\frac{d}{dt} \left(E_{\varepsilon}(\varphi_{p},\varphi_{d}) + \frac{\delta}{2} \|q\|_{L^{2}(\Omega)}^{2} \right) + \sum_{i=p,d} \left(M_{i} \|\nabla\mu_{i}\|_{L^{2}(\Omega)}^{2} + \delta \|\nabla\partial_{t}\varphi_{i}\|_{L^{2}(\Omega)}^{2} \right)
+ \delta \|\Delta q\|_{L^{2}(\Omega)}^{2} + \|\nabla q + T(\varphi_{p})\nabla\mu_{p} + T(\varphi_{d})\nabla\mu_{d}\|_{L^{2}(\Omega)}^{2}
= \int_{\Omega} (S_{p}\mu_{p} + S_{d}\mu_{d}) + (S_{p} + S_{d})q \, dx.$$
(3.30)

Testing now (3.3a) with φ_p , (3.3c) with φ_d and summing the obtained relations yields

$$\frac{1}{2} \frac{d}{dt} \left(\|\varphi_p\|_{L^2(\Omega)}^2 + \|\varphi_d\|_{L^2(\Omega)}^2 \right) \\
= -\sum_{i=p,d} \int_{\Omega} M_i \nabla \mu_i \cdot \nabla \varphi_i - S_i \varphi_i \, dx \\
- \int_{\Omega} \left(\nabla q + T(\varphi_p) \nabla \mu_p + T(\varphi_d) \nabla \mu_d \right) \cdot \left(T(\varphi_p) \nabla \varphi_p + T(\varphi_d) \nabla \varphi_d \right) \, dx.$$
(3.31)

Summing (3.30) and (3.31) then gives

$$\frac{d}{dt} \int_{\Omega} \left(F_{\varepsilon}(\varphi_{p},\varphi_{d}) + F_{1}(\varphi_{p},\varphi_{d}) + \sum_{i=p,d} \frac{1}{2} \left(|\nabla\varphi_{i}|^{2} + |\varphi_{i}|^{2} \right) + \frac{\delta}{2} |q|^{2} \right) dx$$

$$+ \delta \|\Delta q\|_{L^{2}(\Omega)}^{2} + \sum_{i=p,d} \left(\frac{1}{2} M_{i} \|\nabla\mu_{i}\|_{L^{2}(\Omega)}^{2} + \delta \|\nabla\partial_{t}\varphi_{i}\|_{L^{2}(\Omega)}^{2} \right)$$

$$+ \frac{1}{2} \|\nabla q + T(\varphi_{p})\nabla\mu_{p} + T(\varphi_{d})\nabla\mu_{d}\|_{L^{2}(\Omega)}^{2}$$

$$\leq C + C \sum_{i=p,d} \left(\|\varphi_{i}\|_{L^{2}(\Omega)}^{2} + \|\nabla\varphi_{i}\|_{L^{2}(\Omega)}^{2} \right) + \int_{\Omega} S_{p}\mu_{p} + S_{d}\mu_{d} + (S_{p} + S_{d})q \, dx.$$
(3.32)

It remains to control the integral on the right-hand side of (3.32). To handle the pressure term we consider, for a.e. $t \in (0,T)$, the function $N_q(t) \in H^2(\Omega) \cap H^1_0(\Omega)$ as the unique solution to the Poisson problem

$$-\Delta N_q(t) = q(t)$$
 in Ω , $N_q(t) = 0$ on Γ .

As $q(t) \in L^2(\Omega)$, elliptic regularity shows that $||N_q||_{H^2(\Omega)} \leq C_* ||q||_{L^2(\Omega)}$. Furthermore, it can be shown that (see for example [26, §2.2])

$$\langle \partial_t q, N_q \rangle = \frac{1}{2} \frac{d}{dt} \| \nabla N_q \|_{L^2(\Omega)}^2,$$

where $\langle \cdot, \cdot \rangle$ is the duality pairing between $H_0^1(\Omega)$ and $H^{-1}(\Omega)$. We additionally claim that $N_q(0) = 0$. Indeed, as q(0) = 0 from (3.4a), the only solution to the Laplace equation with zero Dirichlet condition is zero. Then, upon testing (3.3g) with N_q leads to

$$\begin{split} &\frac{\delta}{2} \frac{d}{dt} \|\nabla N_q\|_{L^2(\Omega)}^2 + \|q\|_{L^2(\Omega)}^2 + \delta \|\nabla q\|_{L^2(\Omega)}^2 \\ &= \int_{\Omega} (S_p + S_d) N_q - (T(\varphi_p) \nabla \mu_p + T(\varphi_d) \nabla \mu_d) \cdot \nabla N_q \, dx \\ &\leq C \Big(1 + \sum_{i=p,d} \|\varphi_i\|_{L^2(\Omega)} \Big) \|N_q\|_{L^2(\Omega)} + \sum_{i=p,d} \|\nabla \mu_i\|_{L^2(\Omega)} \|\nabla N_q\|_{L^2(\Omega)}, \end{split}$$

where we have also used that $(-\delta\Delta q, \Delta N_q) = \delta(\Delta q, q) = -\delta \|\nabla q\|_{L^2(\Omega)}^2$. Therefore, by Young's and Poincaré's inequalities, as well as the estimate $\|N_q\|_{H^2(\Omega)} \leq C_* \|q\|_{L^2(\Omega)}$, we arrive at the following estimate

$$\frac{\delta}{2} \frac{d}{dt} \|\nabla N_q\|_{L^2(\Omega)}^2 + \frac{1}{2} \|q\|_{L^2(\Omega)}^2 + \delta \|\nabla q\|_{L^2(\Omega)}^2 \\
\leq C \left(1 + \sum_{i=p,d} \left(\|\nabla \mu_i\|_{L^2(\Omega)}^2 + \|\varphi_i\|_{L^2(\Omega)}^2 \right) \right).$$
(3.33)

By virtue of the computations performed in (3.13) we infer that

$$\int_{\Omega} S_{p}\mu_{p} + S_{d}\mu_{d} dx
\leq \frac{M_{p}}{4} \|\nabla\mu_{p}\|_{L^{2}}^{2} + \frac{M_{d}}{4} \|\nabla\mu_{d}\|_{L^{2}(\Omega)}^{2} + C_{\varepsilon} \left(1 + \|\varphi_{p}\|_{L^{2}(\Omega)}^{2} + \|\varphi_{d}\|_{L^{2}(\Omega)}^{2}\right).$$
(3.34)

Then, let κ be a sufficiently small constant such that $\kappa C \leq \frac{1}{4} \min(M_p, M_d)$, where C is the constant on the right-hand side of (3.33), and adding κ times (3.33) to (3.32) yields

$$\frac{d}{dt} \left(\int_{\Omega} (F_{\varepsilon} + F_{1})(\varphi_{p}, \varphi_{d}) dx + \sum_{i=p,d} \frac{1}{2} \|\varphi_{i}\|_{H^{1}(\Omega)}^{2} + \frac{\delta}{2} \left(\|q\|_{L^{2}(\Omega)}^{2} + \kappa \|\nabla N_{q}\|_{L^{2}(\Omega)}^{2} \right) \right) \\
+ \sum_{i=p,d} \left(\frac{1}{4} M_{i} \|\nabla \mu_{i}\|_{L^{2}(\Omega)}^{2} + \delta \|\nabla \partial_{t}\varphi_{i}\|_{L^{2}(\Omega)}^{2} \right) + \frac{\kappa}{4} \|q\|_{L^{2}(\Omega)}^{2} + \delta \kappa \|\nabla q\|_{L^{2}(\Omega)}^{2} \\
+ \delta \|\Delta q\|_{L^{2}(\Omega)}^{2} + \frac{1}{2} \|\nabla q + T(\varphi_{p})\nabla \mu_{p} + T(\varphi_{d})\nabla \mu_{d}\|_{L^{2}(\Omega)}^{2} \\
\leq C_{\varepsilon} + C_{\varepsilon} \sum_{i=p,d} \|\varphi_{i}\|_{H^{1}(\Omega)}^{2},$$
(3.35)

where we have estimated the last term on the right-hand side of (3.32) as follows:

$$\int_{\Omega} (S_p + S_d) q \, dx \le \frac{\kappa}{4} \|q\|_{L^2(\Omega)}^2 + C \left(1 + \|\varphi_p\|_{L^2(\Omega)}^2 + \|\varphi_d\|_{L^2(\Omega)}^2\right).$$

Then, applying Gronwall's inequality to (3.35) yields the following estimates uniform in δ :

$$\begin{aligned} \| (F_{\varepsilon} + F_{1})(\varphi_{p}, \varphi_{d}) \|_{L^{\infty}(0,T;L^{1}(\Omega))} + \| \varphi_{p} \|_{L^{\infty}(0,T;H^{1}(\Omega))} + \| \varphi_{d} \|_{L^{\infty}(0,T;H^{1}(\Omega))} \leq C_{\varepsilon}, \\ \| \nabla \mu_{p} \|_{L^{2}(0,T;L^{2}(\Omega))} + \| \nabla \mu_{d} \|_{L^{2}(0,T;L^{2}(\Omega))} + \| q \|_{L^{2}(0,T;L^{2}(\Omega))} \leq C_{\varepsilon}, \\ \sqrt{\delta} \big(\| q \|_{L^{\infty}(0,T;L^{2}(\Omega))} + \| \nabla \partial_{t}\varphi_{p} \|_{L^{2}(0,T;L^{2}(\Omega))} + \| \nabla \partial_{t}\varphi_{d} \|_{L^{2}(0,T;L^{2}(\Omega))} \big) \leq C_{\varepsilon}, \end{aligned}$$
(3.36)

also thanks to the fact that $q(0) = \Delta q(0) = N_q(0) = 0$ and that

$$\|\varphi_{i,0,\delta}\|_{H^1(\Omega)} \le C \|\varphi_{i,0}\|_{H^1(\Omega)}$$

from (3.6). Then, testing (3.3g) with q and estimating the right-hand side gives

$$\frac{\delta}{2} \frac{d}{dt} \|q\|_{L^{2}(\Omega)}^{2} + \|\nabla q\|_{L^{2}(\Omega)}^{2} + \delta \|\nabla \Delta q\|_{L^{2}(\Omega)}^{2} \\
\leq C \left(1 + \|q\|_{L^{2}(\Omega)}^{2} + \sum_{i=p,d} \|\varphi_{i}\|_{L^{2}(\Omega)}^{2} + \|\nabla \mu_{i}\|_{L^{2}(\Omega)}^{2}\right) + \frac{1}{2} \|\nabla q\|_{L^{2}(\Omega)}^{2}.$$

In light of (3.36), and recalling the initial condition q(0) = 0, we find that

$$\|\nabla q\|_{L^2(0,T;L^2(\Omega))} \le C_{\varepsilon}.$$
(3.37)

Third estimate. Thanks to the Lipschitz regularity of $F_{\varepsilon,i}$ and $F_{1,i}$ for i = p, d, it is easy to see that by (3.36)

$$|(\mu_i)_{\Omega}|^2 \le C_{\varepsilon} (1 + \|\varphi_p\|_{L^2(\Omega)}^2 + \|\varphi_d\|_{L^2(\Omega)}^2) \in L^{\infty}(0,T).$$

Hence, by Poincaré's inequality and (3.36), we deduce

$$\|\mu_p\|_{L^2(0,T;H^1(\Omega))} + \|\mu_d\|_{L^2(0,T;H^1(\Omega))} \le C_{\varepsilon}.$$
(3.38)

Fourth estimate. Testing (3.3b) with $\Delta \varphi_p$, and in light of (3.38) and the Lipschitz regularity of $F_{\varepsilon,p}$ and $F_{1,p}$, we have

$$\frac{1}{2} \|\Delta\varphi_p\|_{L^2(\Omega)}^2 + \frac{d}{dt} \frac{\delta}{2} \|\Delta\varphi_p\|_{L^2(\Omega)}^2 \le C_{\varepsilon} \left(1 + \|\mu_p\|_{L^2(\Omega)}^2 + \|\varphi_p\|_{L^2(\Omega)}^2 + \|\varphi_d\|_{L^2(\Omega)}^2\right).$$
(3.39)

Recalling (3.7) we see that

$$\delta \|\Delta \varphi_{p,0,\delta}\|_{L^2(\Omega)}^2 \le C\delta(1+\delta^{-1}) \|\varphi_{i,0}\|_{H^1(\Omega)}^2 \le C.$$
(3.40)

Thus, integrating (3.39) in time and applying the elliptic estimate

$$||u||_{H^{2}(\Omega)} \leq C (||\Delta u||_{L^{2}(\Omega)} + ||u||_{L^{2}(\Omega)})$$

(holding when u satisfies, as here, no-flux conditions), we obtain

$$\|\varphi_p\|_{L^2(0,T;H^2(\Omega))} + \|\varphi_d\|_{L^2(0,T;H^2(\Omega))} \le C_{\varepsilon},$$

$$\sqrt{\delta} (\|\varphi_p\|_{L^{\infty}(0,T;H^2(\Omega))} + \|\varphi_d\|_{L^{\infty}(0,T;H^2(\Omega))}) \le C_{\varepsilon}.$$
(3.41)

Then, by inspection of (3.3a) we find that

$$\|\partial_t \varphi_p\|_{H^1(\Omega)'} \le C \big(\|\nabla q\|_{L^2(\Omega)} + \|\nabla \mu_p\|_{L^2(\Omega)} + \|\nabla \mu_d\|_{L^2(\Omega)} + \|S_p\|_{L^2(\Omega)} \big),$$

with a similar relation holding for φ_d . Hence, we infer that

$$\|\partial_t \varphi_p\|_{L^2(0,T;H^1(\Omega)')} + \|\partial_t \varphi_d\|_{L^2(0,T;H^1(\Omega)')} \le C_{\varepsilon}.$$
(3.42)

Fifth estimate. Testing (3.3g) with $q - \Delta q \in H_0^1(\Omega)$ and performing standard computations leads to the analogue of (3.23). Then, multiplying both sides of (3.23) by δ and using a Gronwall argument yields

$$\delta \|q\|_{L^{\infty}(0,T;H^{1}(\Omega))} + \delta \|\nabla \Delta q\|_{L^{2}(0,T;L^{2}(\Omega))} \leq C_{\varepsilon},$$

$$\sqrt{\delta} \left(\|\nabla q\|_{L^{2}(0,T;L^{2}(\Omega))} + \|\Delta q\|_{L^{2}(0,T;L^{2}(\Omega))}\right) \leq C_{\varepsilon}.$$
(3.43)

Then, by inspection of (3.3g), and recalling (3.36), (3.37) and (3.43), we infer

$$\delta \|\partial_t q\|_{L^2(0,T;H^{-1}(\Omega))} \leq C \sum_{i=p,d} \left(1 + \|\nabla \mu_i\|_{L^2(0,T;L^2(\Omega))} + \|\varphi_i\|_{L^2(0,T;L^2(\Omega))} \right) + C \|\nabla q\|_{L^2(0,T;L^2(\Omega))} + C \delta \|\nabla \Delta q\|_{L^2(0,T;L^2(\Omega))} + C \leq C_{\varepsilon}.$$
(3.44)

Meanwhile, testing (3.3b) with $-\delta\Delta\partial_t\varphi_p$ and (3.3d) with $-\delta\Delta\partial_t\varphi_d$ we obtain using (3.40)

$$\delta\big(\|\Delta\partial_t\varphi_p\|_{L^2(0,T;L^2(\Omega))} + \|\Delta\partial_t\varphi_d\|_{L^2(0,T;L^2(\Omega))}\big) \le C_{\varepsilon}.$$
(3.45)

On the other hand, testing (3.3a) with $\sqrt{\delta}\partial_t\varphi_p$, we obtain

$$\begin{split} \sqrt{\delta} \|\partial_t \varphi_p\|_{L^2(\Omega)}^2 &\leq C \left(1 + \sum_{i=,p,d} \left(\|\nabla \mu_i\|_{L^2(\Omega)}^2 + \|\varphi_i\|_{L^2(\Omega)}^2 \right) + \|\nabla q\|_{L^2(\Omega)}^2 + \delta \|\nabla \partial_t \varphi_p\|_{L^2(\Omega)}^2 \right) \\ &+ \frac{1}{2} \sqrt{\delta} \|\partial_t \varphi_p\|_{L^2(\Omega)}^2. \end{split}$$

Recalling (3.36) and (3.37), we then deduce that

$$\sqrt{\delta} \|\partial_t \varphi_p\|_{L^2(0,T;L^2(\Omega))}^2 + \sqrt{\delta} \|\partial_t \varphi_d\|_{L^2(0,T;L^2(\Omega))}^2 \le C_{\varepsilon},$$

whence by repeating the same argument on φ_d and by applying elliptic regularity, (3.45) yields

$$\delta \|\varphi_p\|_{H^1(0,T;H^2(\Omega))} + \delta \|\varphi_d\|_{H^1(0,T;H^2(\Omega))} \le C_{\varepsilon}.$$
(3.46)

3.5 Extension to [0, T]

Thanks to the a priori estimates (3.29), (3.36), (3.38), (3.41), (3.42), (3.43), (3.44), (3.45) and (3.46), which have a uniform character with respect to the time variable, we can extend the local solution obtained from Theorem 3.3 up to the full reference interval [0, T]. This can be achieved by means of a standard contradiction argument which we now outline. Suppose there exists a maximal time of existence $T_m \in (0, T]$ for the weak solution ($\varphi_p, \mu_p, \varphi_d, \mu_d, q, n$) to (3.3). To be precise, T_m is defined as the largest time such that ($\varphi_p, \mu_p, \varphi_d, \mu_d, q, n$) exists with the regularity properties specified in the statement of Theorem 3.3. We want to prove that, in fact, $T_m = T$. If this is not the case, repeating the a priori estimates mentioned above (but now working on the maximal time interval $[0, T_m]$), we deduce in particular that

$$\|\varphi_p\|_{C^0([0,T_m];H^2(\Omega))} + \|\varphi_d\|_{C^0([0,T_m];H^2(\Omega))} + \|q\|_{C^0([0,T_m];H^1_0(\Omega))} \le C_{\varepsilon,\delta},$$

where $C_{\varepsilon,\delta}$ is independent of T_m . Note that, to obtain the above bound, we used in particular (3.45) with the continuous embedding $H^1(0,T_m) \subset C^0([0,T_m])$ and (3.43)-(3.44) with the continuous embedding

$$L^{2}(0,T_{m};H^{3}(\Omega)\cap H^{1}_{0}(\Omega))\cap H^{1}(0,T_{m};H^{-1}(\Omega))\subset C^{0}([0,T_{m}];H^{1}_{0}(\Omega)).$$

In particular, the triple $(\varphi_p(t), \varphi_d(t), q(t))$ remains bounded in $H^2(\Omega) \times H^2(\Omega) \times H^1_0(\Omega)$, and actually (strongly) converges in the same space to a limit $(\varphi_p(T_m), \varphi_d(T_m), q(T_m))$, as $t \nearrow T_m$. This allows us to restart the system by taking $\varphi_p(T_m)$, $\varphi_d(T_m)$ and $q(T_m)$ as new "initial" data (note that the other equations of the system have a quasi-static nature; hence they do not involve any initial data). To be precise, we should observe that we performed the fixed point argument by assuming the initial condition q(0) = 0, while we are restarting the argument from $q(T_m) \neq 0$. On the other hand, it is easy to realize that the choice q(0) = 0 was taken just for convenience (indeed, that initial datum will disappear when taking the limit $\delta \to 0$) and the argument still works for any datum in H_0^1 (as is $q(T_m)$). Hence, restarting from T_m we get a new local solution which is defined on an interval of the form $(T_m, T_m + \epsilon)$ for some $\epsilon > 0$ and still enjoys the regularity properties detailed in Theorem 3.3. This contradicts the maximality of T_m . Hence $T_m = T$.

3.6 Passing to the limit $\delta \to 0$

We now pass to the limit $\delta \to 0$ to obtain a weak solution $(\varphi_p^{\varepsilon}, \mu_p^{\varepsilon}, \varphi_d^{\varepsilon}, \mu_d^{\varepsilon}, q^{\varepsilon}, n^{\varepsilon})$ defined over (0, T) to the following problem:

$$\partial_t \varphi_p = M_p \Delta \mu_p + \operatorname{div} \left(T(\varphi_p) \nabla q \right) + \operatorname{div} \left(T(\varphi_p)^2 \nabla \mu_p + T(\varphi_p) T(\varphi_d) \nabla \mu_d \right) + S_p, \quad (3.47a)$$

$$\mu_p = F_{p,p} \left(\varphi_p - \varphi_d \right) + F_{1,p} \left(\varphi_p - \varphi_d \right) - \Delta \varphi_p \quad (3.47b)$$

$$\mu_p = \Gamma_{\varepsilon,p}(\varphi_p, \varphi_d) + \Gamma_{1,p}(\varphi_p, \varphi_d) - \Delta\varphi_p, \qquad (3.476)$$
$$\partial_t \varphi_d = M_d \Delta \mu_d + \operatorname{div}\left(T(\varphi_d) \nabla q\right) + \operatorname{div}\left(T(\varphi_p) T(\varphi_d) \nabla \mu_p + T(\varphi_d)^2 \nabla \mu_d\right) + S_d, \quad (3.47c)$$

$$\mu_d = F_{\varepsilon,d}(\varphi_p, \varphi_d) + F_{1,d}(\varphi_p, \varphi_d) - \Delta\varphi_d, \qquad (3.47d)$$

$$S_p = \Sigma_p(n,\varphi_p,\varphi_d) + m_{pp}\varphi_p + m_{pd}\varphi_d, \qquad (3.47e)$$

$$S_d = \Sigma_d(n, \varphi_p, \varphi_d) + m_{dp}\varphi_p + m_{dd}\varphi_d, \qquad (3.47f)$$

$$0 = \Delta q + \operatorname{div} \left(T(\varphi_p) \nabla \mu_p + T(\varphi_d) \nabla \mu_d \right) + S_p + S_d, \qquad (3.47g)$$

$$0 = -\Delta n + T(\varphi_p)n, \tag{3.47h}$$

furnished with the initial and boundary conditions

$$\varphi_p(0) = \varphi_{p,0}, \quad \varphi_d(0) = \varphi_{d,0} \text{ in } \Omega, \qquad (3.48a)$$

$$M_i \partial_{\boldsymbol{n}} \mu_i + T(\varphi_i) (\nabla q + T(\varphi_p) \nabla \mu_p + T(\varphi_d) \nabla \mu_d) \cdot \boldsymbol{n} = 0, \quad \partial_{\boldsymbol{n}} \varphi_i = 0 \text{ on } \Gamma, \qquad (3.48b)$$

$$n = 1, \quad q = 0 \text{ on } \Gamma.$$
 (3.48c)

Note that in (3.47) the regularized convex part F_{ε} of the potential F is still present.

Theorem 3.4. Let Assumption 2.1 hold, let $\varepsilon \in (0,1)$ and $\delta \in (0,1)$, and let $F_{\varepsilon} : \mathbb{R}^2 \to [0, +\infty)$ be the Moreau-Yosida approximation of F_0 as detailed in Sec. 3. Let also $\varphi_{i,0,\delta} \in H_n^2(\Omega)$ be the unique solution to (3.5). Then, there exists $\delta_0 > 0$ such that for all $\delta < \delta_0$, the weak solution $(\varphi_p^{\delta,\varepsilon}, \mu_p^{\delta,\varepsilon}, \varphi_d^{\delta,\varepsilon}, \eta_d^{\delta,\varepsilon}, n^{\delta,\varepsilon})$ to (3.3) defined on [0,T] and obtained from Theorem 3.3 satisfies the following properties:

there exist functions (φ^ε_p, μ^ε_p, φ^ε_d, μ^ε_d, q^ε, n^ε) such that for i = p, d and any r < ∞ in two dimensions and any r ∈ [1,6) in three dimensions,

$$\begin{split} \varphi_i^{\delta,\varepsilon} &\to \varphi_i^{\varepsilon} \ \text{weakly}^* \ \text{in} \ L^{\infty}(0,T;H^1(\Omega)) \cap L^2(0,T;H^2(\Omega)) \cap H^1(0,T;H^1(\Omega)') \\ \varphi_i^{\delta,\varepsilon} &\to \varphi_i^{\varepsilon} \ \text{strongly} \ \text{in} \ C^0([0,T];L^r(\Omega)) \cap L^2(0,T;W^{1,r}(\Omega)) \ \text{and} \ \text{a.e.} \ \text{in} \ Q, \\ \mu_i^{\delta,\varepsilon} &\to \mu_i^{\varepsilon} \ \text{weakly} \ \text{in} \ L^2(0,T;H^1(\Omega)), \\ q^{\delta,\varepsilon} &\to q^{\varepsilon} \ \text{weakly} \ \text{in} \ L^2(0,T;H^1(\Omega)), \\ n^{\delta,\varepsilon} &\to n^{\varepsilon} \ \text{weakly}^* \ \text{in} \ L^{\infty}(0,T;W^{2,r}(\Omega)) \ \text{and} \ \text{strongly} \ \text{in} \ L^2(0,T;H^1(\Omega)). \end{split}$$

(2) The tuple $(\varphi_p^{\varepsilon}, \mu_p^{\varepsilon}, \varphi_d^{\varepsilon}, \mu_d^{\varepsilon}, q^{\varepsilon}, n^{\varepsilon})$ satisfies equations (3.47b), (3.47d), (3.47e), (3.47f), (3.47h) a.e. in Q, whereas equations (3.47a), (3.47c) and (3.47g) hold for a.e. $t \in (0,T)$ in the following weak sense:

$$0 = \langle \partial_t \varphi_p^{\varepsilon}, \zeta \rangle + \int_{\Omega} \left(M_p \nabla \mu_p^{\varepsilon} + T(\varphi_p^{\varepsilon}) \left(\nabla q^{\varepsilon} + T(\varphi_p^{\varepsilon}) \nabla \mu_p^{\varepsilon} + T(\varphi_d^{\varepsilon}) \nabla \mu_d^{\varepsilon} \right) \right) \cdot \nabla \zeta - S_p \zeta \, dx,$$

$$0 = \langle \partial_t \varphi_d^{\varepsilon}, \zeta \rangle + \int_{\Omega} \left(M_d \nabla \mu_d^{\varepsilon} + T(\varphi_d^{\varepsilon}) \left(\nabla q^{\varepsilon} + T(\varphi_d^{\varepsilon}) \nabla \mu_d^{\varepsilon} + T(\varphi_p^{\varepsilon}) \nabla \mu_p^{\varepsilon} \right) \right) \cdot \nabla \zeta - S_d \zeta \, dx,$$

$$0 = \int_{\Omega} \left(\nabla q^{\varepsilon} + T(\varphi_p^{\varepsilon}) \nabla \mu_p^{\varepsilon} + T(\varphi_d^{\varepsilon}) \nabla \mu_d^{\varepsilon} \right) \cdot \nabla \xi - (S_p + S_d) \xi \, dx$$

for all $\zeta \in H^1(\Omega)$ and $\xi \in H^1_0(\Omega)$. Moreover, $0 \le n^{\varepsilon} \le 1$ a.e. in Q, and $\varphi_i^{\varepsilon}(0) = \varphi_{0,i}$ a.e. in Ω .

Proof. Recalling the estimate (3.6), we immediately infer the following properties of the initial data $(\varphi_{p,0,\delta}, \varphi_{d,0,\delta})$:

$$\begin{split} \|\varphi_{p,0,\delta}\|_{H^1(\Omega)} + \|\varphi_{d,0,\delta}\|_{H^1(\Omega)} &\leq C, \\ \varphi_{p,0,\delta} \to \varphi_{p,0,\delta} \to \varphi_{d,0} \text{ weakly in } H^1(\Omega) \text{ and strongly in } L^2(\Omega). \end{split}$$

Furthermore, this choice of initial data for the regularized system (3.3) implies that the estimate (3.36) is uniform in $\delta \in (0, \delta_0)$.

Then, most of the weak/weak* convergence properties in the statement are directly deduced from the uniform estimates (3.29), (3.36), (3.38) and (3.41), while the strong convergences follow from using [43, §8, Cor. 4]. On the other hand, the strong convergence of $n^{\delta,\varepsilon}$ is proved, similarly as before, by a Cauchy argument which we now sketch. Let (a small) $\eta > 0$ and (a large) $C_* > 0$ be given but otherwise arbitrary. Then, thanks to the a.e. convergence of $\varphi_p^{\delta,\varepsilon}$ to φ_p^{ε} in Q, by Egorov's theorem there exists a measurable subset $X_{\eta} \subset Q$ with $C_*|X_{\eta}| < \frac{1}{4}\eta$ and $\varphi_p^{\delta,\varepsilon} \to \varphi_p^{\varepsilon}$ uniformly in the complement $Q \setminus X_{\eta}$. By this uniform convergence, there exists $\delta_* > 0$ such that for any two indices $0 < \delta_1, \delta_2 < \delta_*$,

$$C_* \int_{Q \setminus X_{\eta}} |T(\varphi_p^{\delta_1,\varepsilon}) - T(\varphi_p^{\delta_2,\varepsilon})|^2 \, dx \, dt < \frac{\eta}{2}.$$

Then, following the computation in (3.28) and using the boundedness of T, we find that

$$\begin{split} \|n^{\delta_{1},\varepsilon} - n^{\delta_{2},\varepsilon}\|_{L^{2}(0,T;H^{1}(\Omega))}^{2} &\leq C_{*}\|T(\varphi_{p}^{\delta_{1},\varepsilon}) - T(\varphi_{p}^{\delta_{2},\varepsilon})\|_{L^{2}(0,T;L^{2}(\Omega))}^{2} \\ &\leq C_{*}\int_{Q\setminus X_{\eta}}|T(\varphi_{p}^{\delta_{1},\varepsilon}) - T(\varphi_{p}^{\delta_{2},\varepsilon})|^{2} \,dx \,dt + C_{*}\int_{X_{\eta}}|T(\varphi_{p}^{\delta_{1},\varepsilon}) - T(\varphi_{p}^{\delta_{2},\varepsilon})|^{2} \,dx \,dt \\ &< \frac{\eta}{2} + 2C_{*}|X_{\eta}| < \eta, \end{split}$$

for $0 < \delta_1, \delta_2 < \delta_*$. Here C_* is exactly the constant C in (3.28). This shows that $\{n^{\delta,\varepsilon}\}_{\delta\in(0,\delta_*)}$ is a Cauchy sequence in $L^2(0,T; H^1(\Omega))$. The property $0 \le n^{\varepsilon} \le 1$ a.e. in Q can be deduced also from a weak comparison principle.

Now passing to the limit $\delta \to 0$ in (3.3e), (3.3f), (3.3h) lead to (3.47e), (3.47f) and (3.47h), respectively. Let us fix $\zeta \in L^2(0,T; H^1(\Omega))$ and test (3.3b) with ζ . Then,

$$\int_0^T \int_\Omega \left(\mu_p^{\delta,\varepsilon} + \Delta \varphi_p^{\delta,\varepsilon} - (F_{\varepsilon,p} + F_{1,p})(\varphi_p^{\delta,\varepsilon}, \varphi_d^{\delta,\varepsilon}) \right) \zeta - \delta \nabla \partial_t \varphi_p^{\delta,\varepsilon} \cdot \nabla \zeta \, dx \, dt = 0$$

Using the weak convergences of $\mu_p^{\delta,\varepsilon}$, $\Delta \varphi_p^{\delta,\varepsilon}$ in $L^2(0,T;L^2(\Omega))$ and the Lipschitz continuity of $F_{\varepsilon,p}$ and $F_{1,p}$, as well as the boundedness $\|\sqrt{\delta}\nabla \partial_t \varphi_p^{\delta,\varepsilon}\|_{L^2(0,T;L^2(\Omega))} \leq C_{\varepsilon}$ resulting from (3.36), passing to the limit $\delta \to 0$ in the above equality leads to

$$\int_0^T \int_\Omega \left(\mu_p^\varepsilon + \Delta \varphi_p^\varepsilon - (F_{\varepsilon,p} + F_{1,p})(\varphi_p^\varepsilon, \varphi_d^\varepsilon) \right) \zeta \, dx \, dt = 0.$$

Since the above identity holds for arbitrary $\zeta \in L^2(0, T; H^1(\Omega))$ and all the terms in the integrand belong to $L^2(0, T; L^2(\Omega))$, the fundamental lemma of calculus of variations then yields (3.47b).

In a similar fashion, we infer from testing (3.3a) with an arbitrary test function $\zeta \in L^2(0,T; H^1(\Omega))$ and then passing to the limit $\delta \to 0$ the identity

$$\begin{split} 0 &= \int_0^T \langle \partial_t \varphi_p^\varepsilon, \zeta \rangle \, dt - \int_0^T \int_\Omega S_p(n^\varepsilon, \varphi_p^\varepsilon, \varphi_d^\varepsilon) \zeta \, dx \, dt \\ &+ \int_0^T \int_\Omega \left(M_p \nabla \mu_p^\varepsilon + T(\varphi_p^\varepsilon) \big(\nabla q^\varepsilon + T(\varphi_p^\varepsilon) \nabla \mu_p^\varepsilon + T(\varphi_d^\varepsilon) \nabla \mu_d^\varepsilon \big) \big) \cdot \nabla \zeta \, dx \, dt, \end{split}$$

where in the above we used the strong L^2 -convergences of $n^{\delta,\varepsilon}$ and $\varphi_i^{\delta,\varepsilon}$ with the generalized Lebesgue dominated convergence theorem and the assumption (2.1) to deduce that $S_p(n^{\delta,\varepsilon}, \varphi_p^{\delta,\varepsilon}, \varphi_d^{\delta,\varepsilon})$ converges to $S_p(n^{\varepsilon}, \varphi_p^{\varepsilon}, \varphi_d^{\varepsilon})$ strongly in $L^2(0, T; L^2(\Omega))$. Furthermore, by the continuity and boundedness of T, it is easy to see that

$$T(\varphi_p^{\delta,\varepsilon}) \to T(\varphi_p^{\varepsilon})$$
 weakly $*$ in $L^{\infty}(Q)$ and strongly in $L^p(Q)$ for all $p \in [1,\infty)$.

Moreover, the strong convergence of the initial data $\varphi_{p,0,\delta}$ to $\varphi_{p,0}$ in $L^2(\Omega)$ and the strong convergence of $\varphi_p^{\delta,\varepsilon}$ to φ_p^{ε} in $C^0([0,T]; L^2(\Omega))$ yield $\varphi_p^{\varepsilon}(0) = \varphi_{p,0}$ as an equality in $L^2(\Omega)$.

Lastly, it remains to pass to the limit in (3.3g). Consider testing (3.3g) with the product $\eta(t)\xi(x)$ for arbitrary test functions $\eta \in C^1(0,T)$ with $\eta(T) = 0$ and $\xi \in H^2(\Omega) \cap H^1_0(\Omega)$, then we have

$$0 = \int_0^T \int_\Omega -\delta q^{\delta,\varepsilon} \xi \partial_t \eta - (S_p + S_d) (n^{\delta,\varepsilon}, \varphi_p^{\delta,\varepsilon}, \varphi_d^{\delta,\varepsilon}) \eta(t) \xi \, dx \, dt \\ + \int_0^T \eta(t) \int_\Omega \left(\nabla q^{\delta,\varepsilon} + T(\varphi_p^{\delta,\varepsilon}) \nabla \mu_p^{\delta,\varepsilon} + T(\varphi_d^{\delta,\varepsilon}) \nabla \mu_d^{\delta,\varepsilon} \right) \cdot \nabla \xi + \delta \Delta q^{\delta,\varepsilon} \cdot \Delta \xi \, dx \, dt.$$

Thanks to $\|q^{\delta,\varepsilon}\|_{L^2(0,T;L^2(\Omega))} \leq C_{\varepsilon}$ from (3.36) and $\sqrt{\delta} \|\Delta q^{\delta,\varepsilon}\|_{L^2(0,T;L^2(\Omega))} \leq C_{\varepsilon}$ from (3.43), after passing to the limit we obtain

$$0 = \int_0^T \eta(t) \int_\Omega \left(\nabla q^\varepsilon + T(\varphi_p^\varepsilon) \nabla \mu_p^\varepsilon + T(\varphi_d^\varepsilon) \nabla \mu_d^\varepsilon \right) \cdot \nabla \xi - (S_p + S_d) (n^\varepsilon, \varphi_p^\varepsilon, \varphi_d^\varepsilon) \xi \, dx \, dt,$$

holding for all $\xi \in H^2(\Omega) \cap H^1_0(\Omega)$ and $\eta \in C^1(0,T)$ with $\eta(T) = 0$. Using the density of $H^2(\Omega) \cap H^1_0(\Omega)$ in $H^1_0(\Omega)$ and the fundamental lemma of calculus of variations, we obtain the weak formulation of (3.47g) as stated in Theorem 3.4.

4 Alternative proof via Galerkin approximation

T

We prove here existence of a solution $(\varphi_p^{\varepsilon}, \mu_p^{\varepsilon}, \varphi_d^{\varepsilon}, \mu_d^{\varepsilon}, q^{\varepsilon}, n^{\varepsilon})$ to (3.47a)-(3.47h) with the initial and boundary conditions (3.48a)-(3.48c) by means of the alternative Faedo-Galerkin argument. This can be stated as follows:

Theorem 4.1. Let Assumption 2.1 hold, let $\varepsilon \in (0,1)$ and let $F_{\varepsilon} : \mathbb{R}^2 \to [0,+\infty)$ be the Moreau-Yosida approximation of F_0 as detailed in Sec. 3. Then, there exists a tuple $(\varphi_p^{\varepsilon}, \mu_p^{\varepsilon}, \varphi_d^{\varepsilon}, \mu_d^{\varepsilon}, q^{\varepsilon}, n^{\varepsilon})$ satisfying assertion (2) of Theorem 3.4. Furthermore, the following energy identity also holds:

$$\frac{d}{dt} \left(\int_{\Omega} (F_{\varepsilon} + F_{1})(\varphi_{p}^{\varepsilon}, \varphi_{d}^{\varepsilon}) + \frac{1}{2} \left(|\nabla \varphi_{p}^{\varepsilon}|^{2} + |\nabla \varphi_{d}^{\varepsilon}|^{2} \right) dx \right)
+ M_{p} ||\nabla \mu_{p}^{\varepsilon}||_{L^{2}(\Omega)}^{2} + M_{d} ||\nabla \mu_{d}^{\varepsilon}||_{L^{2}(\Omega)}^{2} + ||\nabla q^{\varepsilon} + T(\varphi_{p}^{\varepsilon})\nabla \mu_{p}^{\varepsilon} + T(\varphi_{d}^{\varepsilon})\nabla \mu_{d}^{\varepsilon}||_{L^{2}(\Omega)}^{2}$$

$$= \int_{\Omega} (S_{p} + S_{d})q^{\varepsilon} + S_{p}\mu_{p}^{\varepsilon} + S_{d}\mu_{d}^{\varepsilon} dx.$$
(4.1)

Proof. Let $\{w_i\}_{i\in\mathbb{N}}$ and $\{y_i\}_{i\in\mathbb{N}}$ denote the eigenfunctions of the Neumann-Laplacian operator $-\Delta + I$ and of the Dirichlet-Laplacian operator $-\Delta$, respectively. It is then well-known that $\{w_i\}_{i\in\mathbb{N}}$ and $\{y_i\}_{i\in\mathbb{N}}$ may be normalized in such a way to form orthogonal bases of $H^1(\Omega)$ and $H^1_0(\Omega)$, respectively, that are also orthonormal with respect to the $L^2(\Omega)$ -scalar product. Furthermore, elliptic regularity then yields that $w_i \in H^2_n(\Omega)$ and $y_i \in H^2(\Omega) \cap H^1_0(\Omega)$ for all $i \in \mathbb{N}$.

For fixed $k \in \mathbb{N}$, we seek functions $\varphi_{p,k}, \mu_{p,k}, \varphi_{d,k}, \mu_{d,k} \in \operatorname{span}\{w_1, \ldots, w_k\}, q_k, n_k - 1 \in \operatorname{span}\{y_1, \ldots, y_k\}$ which solve for all $1 \leq j \leq k$ and $i \in \{p, d\}$:

$$0 = (\varphi'_{i,k}, w_j) + (M_i \nabla \mu_{i,k}, \nabla w_j) - (T(\varphi_{i,k}) u_k, \nabla w_j) - (S_{i,k}, w_j),$$
(4.2a)

$$0 = (\mu_{i,k}, w_j) - (\nabla \varphi_{i,k}, \nabla w_j) - ((F_{\varepsilon,i} + F_{1,i})(\varphi_{p,k}, \varphi_{d,k}), w_k),$$

$$(4.2b)$$

$$\boldsymbol{u}_{k} = -\nabla q_{k} - T(\varphi_{p,k})\nabla \mu_{p,k} - T(\varphi_{d,k})\nabla \mu_{d,k}, \qquad (4.2c)$$

$$0 = (u_k, \nabla y_j) + (S_{p,k} + S_{d,k}, y_j),$$
(4.2d)

$$S_{i,k} = S_i(n_k, \varphi_{p,k}, \varphi_{d,k}), \tag{4.2e}$$

$$0 = (\nabla n_k, \nabla y_j) + (T(\varphi_{p,k})n_k, y_j), \tag{4.2f}$$

furnished with the initial condition $\varphi_{i,k}(0) = \prod_k(\varphi_{i,0})$, where \prod_k denotes the projection to the finite dimension subspace span $\{w_1, \ldots, w_k\}$ and $\varphi'_{i,k} = \frac{d}{dt}\varphi_{i,k}$. Moreover, (\cdot, \cdot) denotes the usual scalar product in $L^2(\Omega)$. We claim that the above system (4.2) admits a (local in time) solution. Indeed, we have

$$\varphi_{i,k}(t) = \sum_{j=1}^{k} \alpha_{i,j}^{k}(t) w_{j}, \quad \mu_{i,k}(t) = \sum_{j=1}^{k} \beta_{i,j}^{k}(t) w_{j},$$
$$q_{k}(t) = \sum_{j=1}^{k} \gamma_{j}^{k}(t) y_{j}, \quad n_{k}(t) = 1 + \sum_{j=1}^{k} \eta_{j}^{k}(t) y_{j},$$

and we set $\boldsymbol{\alpha}_{i}^{k} := (\alpha_{i,1}^{k}, \dots, \alpha_{i,k}^{k}), \ \boldsymbol{\beta}_{i}^{k} := (\beta_{i,1}^{k}, \dots, \beta_{i,k}^{k}), \ i = p, d, \ \boldsymbol{\gamma}^{k} := (\gamma_{1}^{k}, \dots, \gamma_{k}^{k})$ and $\boldsymbol{\eta}^{k} := (\eta_{1}^{k}, \dots, \eta_{k}^{k})$. Then, from (4.2b) the $\boldsymbol{\beta}_{i}^{k}$ can be immediately expressed as globally Lipschitz continuous functions of the $\boldsymbol{\alpha}_{i}^{k}$ (i = p, d). From (4.2d), taking (4.2c) into account, we can also express the $\boldsymbol{\gamma}^{k}$ as globally Lipschitz continuous functions of the $\boldsymbol{\alpha}_{i}^{k}$ (i = p, d). From (4.2d), taking (4.2c) into account, we can also express the $\boldsymbol{\gamma}^{k}$ as globally Lipschitz continuous functions of the $\boldsymbol{\alpha}_{i}^{k}$ (i = p, d) and $\boldsymbol{\eta}^{k}$. It is now possible to express the $\boldsymbol{\eta}^{k}$ in terms of the $\boldsymbol{\alpha}_{p}^{k}$ via the equation (4.2f) (see [28, §6]). Indeed, (4.2f) leads to the following algebraic system

$$\lambda_{\ell}\eta_{\ell}^{k} + \sum_{j=1}^{k} \left(T(\varphi_{p,k})y_{j}, y_{\ell} \right) \eta_{j}^{k} = -\left(T(\varphi_{p,k}), y_{\ell} \right), \qquad \ell = 1, \cdots k, \qquad (4.3)$$

where λ_{ℓ} are the eigenvalues of the Dirichlet-Laplacian operator. We now check that the matrix $\mathbb{A} := (a_{jl})_{j,l=1...k}$ of entries $a_{jl} := \lambda_{\ell} \delta_{jl} + (T(\varphi_{p,k})y_l, y_j)$ is invertible. To this aim,

it is enough to prove that A is positive definite. Take $\boldsymbol{\zeta} := (\zeta_1, \ldots, \zeta_k) \in \mathbb{R}^k$ and set $\phi := \sum_{j=1}^k \zeta_j y_j$. We have

$$\sum_{j,l=1}^{k} a_{jl} \zeta_j \zeta_l = \|\nabla \phi\|_{L^2(\Omega)}^2 + \left(T(\varphi_{p,k})\phi,\phi\right) \ge \|\nabla \phi\|_{L^2(\Omega)}^2 \ge \lambda_1 \|\phi\|_{L^2(\Omega)}^2.$$
(4.4)

Therefore, since the y_j , j = 1, ..., k, are linearly independent, we have $\sum_{j,l=1}^k a_{jl} \zeta_j \zeta_l = 0$ iff $\phi = 0$ iff $\zeta = 0$, and this implies that \mathbb{A} is positive definite and hence invertible, for each choice of the α_p^k . Hence, by inverting system (4.3) we can express the η^k as globally Lipschitz continuous functions of the α_p^k . This also entails that the γ^k are globally Lipschitz continuous functions of the α_i^k only. From (4.2a), taking also (4.2c) into account, we are led to an ODE system of 2k equations in the 2k unknowns $\alpha_{i,j}^k$ (i = p, d) in normal form, with some initial conditions that can be deduced from $\varphi_{i,k}(0) = \Pi_k(\varphi_{i,0})$. The Cauchy-Lipschitz theorem can be applied to this system and we can therefore guarantee the existence of its unique maximal solution $\alpha_i^k \in C^1([0, T_k^*); \mathbb{R}^k)$, for some $T_k^* \in (0, \infty]$. Finally, β_i^k , $i = p, d, \gamma^k$ and η^k are obtained as well. We have thus proven that the finitedimensional problem (4.2a)–(4.2f), endowed with the initial condition $\varphi_{i,k}(0) = \Pi_k(\varphi_{i,0})$, has a (unique) maximal solution $\alpha_i^k, \beta_i^k, \gamma^k, \eta^k \in C^1([0, T_k^*); \mathbb{R}^k)$.

We now derive estimates that are uniform in k. Multiplying (4.2a) with $\beta_{i,j}^k$, (4.2b) with $\alpha_{i,j}^k$, (4.2c) with u_k , (4.2a) with $\alpha_{i,j}^k$, upon summing the ensuing identities (over $j = 1 \dots k$, and i = p, d) and arguing as for (3.30), (3.31), we are led to an analogous inequality to (3.32) with δ set to be zero. The source term involving $S_{p,k}\mu_{p,k} + S_{d,k}\mu_{d,k}$ can be handled as in (3.34), as due to the sublinear growth of ∇F_{ε} , ∇F_1 one obtains that

$$|(\mu_{i,k})_{\Omega}| \le C_{\varepsilon} \left(1 + \|\varphi_{p,k}\|_{L^{2}(\Omega)} + \|\varphi_{d,k}\|_{L^{2}(\Omega)} \right).$$
(4.5)

Meanwhile, for the source term involving $(S_{p,k} + S_{d,k})q_k$, we observe from (4.2e) that

$$||S_{p,k}||_{L^{2}(\Omega)} + ||S_{d,k}||_{L^{2}(\Omega)} \le C(1 + ||\varphi_{p,k}||_{L^{2}(\Omega)} + ||\varphi_{d,k}||_{L^{2}(\Omega)}).$$

Testing (4.2c) with ∇q_k and (4.2d) with γ_j^k , summing over $j = 1 \dots k$ and combining the resulting identities yields

$$\|\nabla q_k\|_{L^2(\Omega)}^2 \le C \|S_{p,k} + S_{d,k}\|_{L^2(\Omega)} \|q_k\|_{L^2(\Omega)} + \left(\|\nabla \mu_{p,k}\|_{L^2(\Omega)} + \|\nabla \mu_{d,k}\|_{L^2(\Omega)}\right) \|\nabla q_k\|_{L^2(\Omega)}.$$

Young's inequality and Poincaré's inequality then give

$$\|q_k\|_{H^1(\Omega)} \le C \left(1 + \|\varphi_{p,k}\|_{L^2(\Omega)} + \|\varphi_{d,k}\|_{L^2(\Omega)} + \|\nabla\mu_{p,k}\|_{L^2(\Omega)} + \|\nabla\mu_{d,k}\|_{L^2(\Omega)} \right).$$
(4.6)

A further application of Young's inequality entails

$$\int_{\Omega} \left| (S_{p,k} + S_{d,k}) q_k \right| dx \leq \frac{M_p}{4} \| \nabla \mu_{p,k} \|_{L^2(\Omega)}^2 + \frac{M_d}{4} \| \nabla \mu_{d,k} \|_{L^2(\Omega)}^2 + C \left(1 + \| \varphi_{p,k} \|_{L^2(\Omega)}^2 + \| \varphi_{d,k} \|_{L^2(\Omega)}^2 \right),$$

and consequently we obtain the analogue to (3.35) with δ and κ set to zero. In particular, by employing (4.5) and the Poincaré inequality to control the sequence of $\mu_{i,k}$ in $L^2(0,T; L^2(\Omega))$, and also using the convergences $\varphi_{i,k}(0) \to \varphi_{i,0}$ in $H^1(\Omega)$ as $k \to \infty$, for i = p, d (cf. (A4)), we first get $T_k^* = +\infty$ (notice that $\|\varphi_{i,k}(t)\| = |\alpha_i^k(t)|$), and then we obtain the following uniform estimate which holds for any given $0 < T < +\infty$,

$$\| (F_{\varepsilon} + F_1)(\varphi_{p,k}, \varphi_{d,k}) \|_{L^{\infty}(0,T;L^1(\Omega))} + \| \varphi_{i,k} \|_{L^{\infty}(0,T;H^1(\Omega))} + \| \mu_{i,k} \|_{L^2(0,T;H^1(\Omega))} + \| u_k \|_{L^2(0,T;L^2(\Omega))} \le C_{\varepsilon} .$$

$$(4.7)$$

Taking the L^2 -norm of ∇q_k in (4.2c) and using the above estimates for $\boldsymbol{u}_k, \mu_{i,k}$ yields

$$\|q_k\|_{L^2(0,T;H^1_0(\Omega))} \le C.$$
(4.8)

Using the sublinear growth of ∇F_{ε} , ∇F_1 again, together with (4.7), we get also

$$\|(F_{\varepsilon,i} + F_{1,i})(\varphi_{p,k}, \varphi_{d,k})\|_{L^{\infty}(0,T;L^{2}(\Omega))} \leq C, \qquad i = p, d.$$
(4.9)

Testing (4.2b) with the coefficients of $\Delta \varphi_{i,k}$, and using the sublinear growth of ∇F_1 , as well as the convexity of F_{ε} , by means of a standard argument we infer the boundedness of $\Delta \varphi_{i,k}$ in $L^2(0,T;L^2(\Omega))$. Elliptic regularity then yields

$$\|\varphi_{i,k}\|_{L^2(0,T;H^2(\Omega))} \le C_{\varepsilon}.$$

$$(4.10)$$

Moreover, by comparison in (4.2a), and by relying on the basic estimates (4.7), a standard argument entails also a bound for the sequences of the time derivatives $\varphi'_{p,k}$, $\varphi'_{d,k}$, namely

$$\|\varphi_{i,k}'\|_{L^2(0,T;(H^1(\Omega))')} \le C_{\varepsilon}.$$
(4.11)

Concerning the estimate for the sequence of n_k , testing (4.2f) with η_j^k and summing over $j = 1 \dots k$, we get

$$\|\nabla n_k\|_{L^2(\Omega)}^2 + \left(T(\varphi_{p,n})n_k, n_k\right) = \left(T(\varphi_{p,n}), n_k\right).$$

Using the boundedness and the nonnegativity of $T(\cdot)$ we immediately infer the bound

$$\|n_k\|_{L^{\infty}(0,T;H^1(\Omega))} \le C.$$
(4.12)

Thanks to estimates (4.7)–(4.12), and to Aubin-Lions lemma, we deduce there exist φ_p , φ_d , μ_p , μ_d , \boldsymbol{u} , q and n in the regularity class

$$\varphi_p, \varphi_d \in L^2(0, T; H^2(\Omega)) \cap L^{\infty}(0, T; H^1(\Omega)) \cap H^1(0, T; H^1(\Omega)'),$$
 (4.13)

$$\mu_p, \mu_d \in L^2(0, T; H^1(\Omega)), \qquad (4.14)$$

$$\boldsymbol{u} \in L^2(0, T; L^2(\Omega)^3),$$
 (4.15)

$$q \in L^2(0, T; H^1_0(\Omega)), \qquad (4.16)$$

$$n \in 1 + L^{\infty}(0, T; H_0^1(\Omega)),$$
(4.17)

such that, for a not relabelled subsequence, we have

$$\varphi_{i,k} \to \varphi_i \quad \text{weakly} * \text{ in } L^2(0,T;H^2(\Omega)) \cap H^1(0,T;H^1(\Omega)') \cap L^\infty(0,T;H^1(\Omega)), \quad (4.18)$$

$$\varphi_{i,k} \to \varphi_i \quad \text{strongly in } L^2(0,T; H^{2-\sigma}(\Omega)) \quad (\sigma > 0), \text{ and a.e. in } Q_T,$$

$$(4.19)$$

$$\mu_{i,k} \to \mu_i \quad \text{weakly in } L^2(0,T; H^1(\Omega)),$$

$$(4.20)$$

$$\boldsymbol{u}_k \to \boldsymbol{u} \quad \text{weakly in } L^2(0,T;L^2(\Omega)^3),$$

$$(4.21)$$

$$q_k \to q \quad \text{weakly in } L^2(0,T;H^1_0(\Omega)),$$

$$(4.22)$$

$$n_k \to n \quad \text{weakly}^* \text{ in } L^{\infty}(0,T; H^1(\Omega)),$$

$$(4.23)$$

Moreover, thanks to the continuity of $F_{\varepsilon,i} + F_{1,i}$ and to (4.9), (4.19), we also have

$$(F_{\varepsilon,i} + F_{1,i})(\varphi_{p,k}, \varphi_{d,k}) \to (F_{\varepsilon,i} + F_{1,i})(\varphi_p, \varphi_d) \quad \text{weakly} * \text{ in } L^{\infty}(0,T; L^2(\Omega)).$$
(4.24)

We now need to derive strong convergence for the sequence of n_k (or at least for a subsequence), in order to pass to the limit in (4.2e), by showing that $\{n_k\}_{k\in\mathbb{N}}$ is a Cauchy sequence in $L^2(0,T; H^1(\Omega))$. Here we cannot directly adapt the argument of the proof of Theorem 3.4, since here a uniform L^{∞} bound for the Galerkin approximants n_k seems not to be available from a weak comparison principle. However, the argument is still simple. Indeed, after writing (4.2f) for two different indexes k and ℓ , taking the difference, multiplying the resulting identity by $\eta_i^k - \eta_i^\ell$ and summing over $j = 1 \dots k$, we obtain

$$\|\nabla(n_k - n_\ell)\|_{L^2(\Omega)}^2 + \left(T(\varphi_{p,k})(n_k - n_\ell), n_k - n_\ell\right) = -\left((T(\varphi_{p,k}) - T(\varphi_{p,\ell}))n_\ell, n_k - n_\ell\right).$$

Thanks to the nonnegativity of $T(\cdot)$ and to the Poincaré inequality, then there follows that

$$\|\nabla(n_k - n_\ell)\|_{L^2(\Omega)}^2 \le C \|(T(\varphi_{p,k}) - T(\varphi_{p,\ell}))n_\ell\|_{L^2(\Omega)}^2 \le C \|\varphi_{p,k} - \varphi_{p,\ell}\|_{L^\infty(\Omega)}^2 \|n_\ell\|_{L^2(\Omega)}^2.$$

We now use the strong convergence (4.19), which implies that $\{\varphi_{p,k}\}$ is a Cauchy sequence in $L^2(0,T;L^{\infty}(\Omega))$ (by taking $\sigma > 0$ small enough), to deduce that $\{n_k\}$ is a Cauchy sequence in $L^2(0,T;H_0^1(\Omega))$ and hence we get

$$n_k \to n \quad \text{strongly in } L^2(0,T;H^1(\Omega)).$$

$$(4.25)$$

The weak/strong convergences (4.18)–(4.25) are now enough to pass to the limit in the approximate problem (4.2a)–(4.2f) by means of a standard argument and prove that the tuple $(\varphi_p, \mu_p, \varphi_d, \mu_d, q, n)$ satisfies assertion (2) of Theorem 3.4. In particular, from (4.2f) we obtain in the limit that $(\nabla n, \nabla y) + (T(\varphi_p)n, y) = 0$ for all $y \in H_0^1(\Omega)$ which is the weak form of (1.6g). Then, the weak comparison principle and elliptic regularity can now be applied, leading, respectively, to the pointwise bound $0 \le n \le 1$ a.e in $\Omega \times (0, T)$ and to the further regularity $n \in L^{\infty}(0,T; W^{2,r}(\Omega))$ for every $r < \infty$.

To show that (4.1) holds for the weak solution, it suffices to show that we have sufficient regularity to test the equation for φ_i^{ε} with μ_i^{ε} , the equation for μ_i^{ε} with $\partial_t \varphi_i^{\varepsilon}$, the equation for $\boldsymbol{u}^{\varepsilon}$ with $\boldsymbol{u}^{\varepsilon}$, to sum the resulting relations and to integrate by parts. This is actually possible, and the main technical detail lies in showing that

$$\sum_{i=p,d} \langle \partial_t \varphi_i^{\varepsilon}, \mu_i^{\varepsilon} \rangle = \frac{d}{dt} \int_{\Omega} (F_{\varepsilon} + F_1)(\varphi_p^{\varepsilon}, \varphi_d^{\varepsilon}) + \frac{1}{2} (|\nabla \varphi_p^{\varepsilon}|^2 + |\nabla \varphi_d^{\varepsilon}|^2) \, dx.$$
(4.26)

To see this, we define the convex and lower semicontinuous functional $\mathcal{G}: L^2(\Omega) \times L^2(\Omega) \to \mathbb{R} \cup \{+\infty\}$ by

$$\mathcal{G}(\varphi_p,\varphi_d) := \begin{cases} \int_{\Omega} F_{\varepsilon}(\varphi_p,\varphi_d) + \frac{1}{2} (|\nabla \varphi_p|^2 + |\nabla \varphi_d|^2) \, dx & \text{if } (\varphi_p,\varphi_d) \in (H^1(\Omega))^2 \\ & \text{and } F_{\varepsilon}(\varphi_p,\varphi_d) \in L^1(\Omega), \\ +\infty & \text{otherwise.} \end{cases}$$

Then, by [2, Proposition 2.8] the subdifferential of \mathcal{G} can be characterized as

$$\partial \mathcal{G}(\varphi_p, \varphi_d) = (-\Delta \varphi_p + F_{\varepsilon, p}(\varphi_p, \varphi_d), -\Delta \varphi_d + F_{\varepsilon, d}(\varphi_p, \varphi_d))$$

for all $(\varphi_p, \varphi_d) \in D(\partial \mathcal{G}) = (H_n^2(\Omega))^2$. Moreover, thanks to the Lipschitz continuity of F_1 , we have

$$\sum_{i=p,d} \langle \partial_t \varphi_i^{\varepsilon}, F_{1,i}(\varphi_p^{\varepsilon}, \varphi_d^{\varepsilon}) \rangle = \frac{d}{dt} \int_{\Omega} F_1(\varphi_p^{\varepsilon}, \varphi_d^{\varepsilon}) \, dx, \tag{4.27}$$

whereas applying standard chain rule formulas for monotone operators in Hilbert spaces (see [6, Lemme 3.3, p. 73], cf. also [16, Proposition 4.2]), we deduce that

$$\langle (\partial_t \varphi_p^\varepsilon, \partial_t \varphi_d^\varepsilon), \partial \mathcal{G}(\varphi_p^\varepsilon, \varphi_d^\varepsilon) \rangle = \frac{d}{dt} \mathcal{G}(\varphi_p^\varepsilon, \varphi_d^\varepsilon).$$

Using that $\mu_i^{\varepsilon} = -\Delta \varphi_i^{\varepsilon} + (F_{\varepsilon,i} + F_{1,i})(\varphi_p^{\varepsilon}, \varphi_d^{\varepsilon})$ holds a.e. in $\Omega \times (0, T)$, the required assertion (4.26) is proved.

Remark 4.1. It is worth observing that the Faedo-Galerkin approximants $\mu_{i,k}$, being linear combinations of the Neumann eigenfunctions $\{w_j\}$, satisfy a no-flux condition on Γ , which is different with respect to the coupled condition expected to hold in the limit (i.e., the first of (1.9)). On the other hand, this is not a contradiction in view of the fact that in the limit $k \to \infty$ we will recover the first of (1.9) (in an implicit form) from the limit of equations (4.2a). Indeed, $\mu_{i,k}$ converges to μ_i in $H^1(\Omega)$, but not in $H^2(\Omega)$ (cf. (4.20)). Then, as $H^2_n(\Omega)$ is dense in $H^1(\Omega)$, the no-flux condition may be (and in fact is) lost in the limit and replaced by a different one.

5 Passing to the limit $\varepsilon \to 0$

Let $(\varphi_p^{\varepsilon}, \mu_p^{\varepsilon}, \varphi_d^{\varepsilon}, \mu_d^{\varepsilon}, q^{\varepsilon}, n^{\varepsilon})$ denote a weak solution to (3.47) obtained either from Theorem 3.4 or from Theorem 4.1. Introducing the velocity variable as $\boldsymbol{u}^{\varepsilon} := -\nabla q^{\varepsilon} - T(\varphi_p^{\varepsilon})\nabla \mu_p^{\varepsilon} - T(\varphi_d^{\varepsilon})\nabla \mu_d^{\varepsilon}$, we can now rewrite (3.47) as

$$\partial_t \varphi_p^{\varepsilon} = M_p \Delta \mu_p^{\varepsilon} - \operatorname{div} \left(T(\varphi_p^{\varepsilon}) \boldsymbol{u}^{\varepsilon} \right) + S_p, \tag{5.1a}$$

$$\mu_p^{\varepsilon} = F_{\varepsilon,p}(\varphi_p^{\varepsilon}, \varphi_d^{\varepsilon}) + F_{1,p}(\varphi_p^{\varepsilon}, \varphi_d^{\varepsilon}) - \Delta \varphi_p^{\varepsilon},$$
(5.1b)

$$\partial_t \varphi_d^{\varepsilon} = M_d \Delta \mu_d^{\varepsilon} - \operatorname{div} \left(T(\varphi_d^{\varepsilon}) \boldsymbol{u}^{\varepsilon} \right) + S_d, \tag{5.1c}$$

$$\mu_d^{\varepsilon} = F_{\varepsilon,d}(\varphi_p^{\varepsilon}, \varphi_d^{\varepsilon}) + F_{1,d}(\varphi_p^{\varepsilon}, \varphi_d^{\varepsilon}) - \Delta \varphi_d^{\varepsilon}, \tag{5.1d}$$

$$S_p = \Sigma_p(n^{\varepsilon}, \varphi_p^{\varepsilon}, \varphi_d^{\varepsilon}) + m_{pp}\varphi_p^{\varepsilon} + m_{pd}\varphi_d^{\varepsilon}, \qquad (5.1e)$$

$$S_d = \Sigma_d(n^\varepsilon, \varphi_p^\varepsilon, \varphi_d^\varepsilon) + m_{dp}\varphi_p^\varepsilon + m_{dd}\varphi_d^\varepsilon, \qquad (5.1f)$$

$$\boldsymbol{u}^{\varepsilon} = -\nabla q^{\varepsilon} - T(\varphi_{p}^{\varepsilon})\nabla \mu_{p}^{\varepsilon} - T(\varphi_{d}^{\varepsilon})\nabla \mu_{d}^{\varepsilon}, \qquad (5.1g)$$

$$\operatorname{div} \boldsymbol{u}^{\varepsilon} = S_p + S_d, \tag{5.1h}$$

$$0 = -\Delta n^{\varepsilon} + T(\varphi_p^{\varepsilon}) n^{\varepsilon}, \qquad (5.1i)$$

furnished with the initial-boundary conditions (3.48a)-(3.48c) (in fact, the system is satisfied in the weak form specified in the statement; nevertheless, it is probably clearer to report the equations in their strong formulation).

The aim of this section is to derive uniform a priori estimates in ε and then pass to the limit $\varepsilon \to 0$. Let us point out that the estimate (3.29) involving n^{ε} is already uniform in ε . For convenience, we will drop the superscript ε in the variables, and denote with the symbol C positive constants that are independent of ε .

5.1 A priori estimates

We will now derive a number of estimates uniform with respect to ε . We start controlling the mean values of φ_p and φ_d . Denoting

$$\boldsymbol{y}(t) := ((\varphi_p)_{\Omega}(t), (\varphi_d)_{\Omega}(t)), \quad (\boldsymbol{\Sigma})_{\Omega} = ((\Sigma_p)_{\Omega}, (\Sigma_d)_{\Omega}),$$

then by testing (5.1a) and (5.1c) with 1 leads to the following system of ODE's:

$$\frac{d}{dt}\boldsymbol{y}(t) = (\boldsymbol{\Sigma})_{\Omega}(t) + \underline{M}\boldsymbol{y}(t)$$
(5.2)

for any $0 \leq t \leq T$. Thanks to (2.1), (2.2) and (2.8) we infer that the vector $\boldsymbol{y}(t) = ((\varphi_p)_{\Omega}(t), (\varphi_d)_{\Omega}(t))$ belongs to the interior int Δ_0 for all times $t \in [0, T]$. Indeed, at the time t = 0, $\boldsymbol{y}(0) \in \text{int } \Delta_0$ by (2.8). Suppose that there exists a time t_* such that $\boldsymbol{y}(t_*) \in \partial \Delta_0$. Then, taking $t = t_*$ in the above ODE, multiplying with the outer unit normal \boldsymbol{n} to Δ_0 and applying (2.2), we necessarily have that

$$\frac{d}{dt}\boldsymbol{y}(t_*)\cdot\boldsymbol{n}<0.$$

As a consequence, $\boldsymbol{y}(t) \in \operatorname{int} \Delta_0$ for t in a right neighbourhood of t_* , whence it is apparent that $\boldsymbol{y}(t)$ can never leave Δ_0 . From this we deduce that there exist positive constants $0 < c_1 < c_2 < 1$ independent of ε such that

$$c_1 \le (\varphi_p)_{\Omega}(t), (\varphi_d)_{\Omega}(t) \le c_2, \quad c_1 \le (\varphi_p + \varphi_d)_{\Omega}(t) \le c_2 \quad \forall t \in [0, T].$$
(5.3)

Testing now (5.1a) with μ_p , (5.1c) with μ_d , (5.1b) with $\partial_t \varphi_p$, (5.1d) with $\partial_t \varphi_d$, (5.1g) with \boldsymbol{u} and summing leads to

$$\frac{d}{dt} \int_{\Omega} F_{\varepsilon}(\varphi_p, \varphi_d) + F_1(\varphi_p, \varphi_d) + \frac{1}{2} \left(|\nabla \varphi_p|^2 + |\nabla \varphi_d|^2 \right) dx
+ M_p ||\nabla \mu_p||^2_{L^2(\Omega)} + M_d ||\nabla \mu_d||^2_{L^2(\Omega)} + ||\boldsymbol{u}||^2_{L^2(\Omega)}
= \int_{\Omega} S_p \mu_p + S_d \mu_d + q(S_p + S_d) dx.$$
(5.4)

In the above we used Darcy's law and integration by parts to deduce that

$$\int_{\Omega} \left(T(\varphi_p) \nabla \mu_p + T(\varphi_d) \nabla \mu_d \right) \cdot \boldsymbol{u} \, d\boldsymbol{x} = \int_{\Omega} -\nabla q \cdot \boldsymbol{u} - |\boldsymbol{u}|^2 \, d\boldsymbol{x} = \int_{\Omega} q(S_p + S_d) - |\boldsymbol{u}|^2 \, d\boldsymbol{x}.$$

Let us now observe that, by the boundedness of Σ_p , we have

$$\begin{split} \int_{\Omega} S_p \mu_p \, dx &\leq C \|\mu_p - (\mu_p)_{\Omega}\|_{L^1(\Omega)} + C |(\mu_p)_{\Omega}| + \sum_{i=p,d} \int_{\Omega} m_{pi} \varphi_i (\mu_p - (\mu_p)_{\Omega} + (\mu_p)_{\Omega}) \, dx \\ &\leq C \|\mu_p - (\mu_p)_{\Omega}\|_{L^1(\Omega)} + C |(\mu_p)_{\Omega}| + \sum_{i=p,d} \int_{\Omega} m_{pi} (\varphi_i - (\varphi_i)_{\Omega}) (\mu_p - (\mu_p)_{\Omega}) \, dx, \\ &+ (\mu_p)_{\Omega} \int_{\Omega} m_{pp} \varphi_p + m_{pd} \varphi_d \, dx \\ &\leq C \|\mu_p - (\mu_p)_{\Omega}\|_{L^1(\Omega)} + C |(\mu_p)_{\Omega}| + C \sum_{i=p,d} \|\nabla \varphi_i\|_{L^2(\Omega)} \|\nabla \mu_p\|_{L^2(\Omega)}, \end{split}$$

where we have used that $((\varphi_p)_{\Omega}, (\varphi_d)_{\Omega})$ never leaves the set Δ_0 and so $m_{pp}(\varphi_p)_{\Omega} + m_{pd}(\varphi_d)_{\Omega}$ is bounded. An analogous estimate holds for $S_d\mu_d$, whence, by the Poincaré and Young inequalities, we obtain

$$\left| \int_{\Omega} S_{p} \mu_{p} + S_{d} \mu_{d} \right| \leq C \left(|(\mu_{p})_{\Omega}| + |(\mu_{d})_{\Omega}| \right) + \frac{M_{p}}{4} \| \nabla \mu_{p} \|_{L^{2}(\Omega)}^{2} + \frac{M_{d}}{4} \| \nabla \mu_{d} \|_{L^{2}(\Omega)}^{2} + C \left(1 + \| \nabla \varphi_{p} \|_{L^{2}(\Omega)}^{2} + \| \nabla \varphi_{d} \|_{L^{2}(\Omega)}^{2} \right).$$
(5.5)

For the term involving the pressure q, we have

$$\int_{\Omega} (S_p + S_d) q \, dx \bigg| \le C_{\eta} \big(1 + \|\varphi_p\|_{L^2(\Omega)}^2 + \|\varphi_d\|_{L^2(\Omega)}^2 \big) + \eta \|q\|_{L^2(\Omega)}^2$$

for some positive constant η to be fixed below. To get an L^2 -estimate of the pressure, we use the Poincaré inequality for $H_0^1(\Omega)$ -functions and Darcy's law to deduce that

$$\|q\|_{L^{2}(\Omega)}^{2} \leq C \|\nabla q\|_{L^{2}(\Omega)}^{2} \leq C \left(\|\boldsymbol{u}\|_{L^{2}(\Omega)}^{2} + \|\nabla \mu_{p}\|_{L^{2}(\Omega)}^{2} + \|\nabla \mu_{d}\|_{L^{2}(\Omega)}^{2}\right).$$
(5.6)

Take now η sufficiently small so that

$$\left| \int_{\Omega} (S_p + S_d) q \, dx \right| \leq \frac{1}{2} \| \boldsymbol{u} \|_{L^2(\Omega)}^2 + \frac{M_p}{4} \| \nabla \mu_p \|_{L^2(\Omega)}^2 + \frac{M_d}{4} \| \nabla \mu_d \|_{L^2(\Omega)}^2 + C \big(1 + \| \varphi_p \|_{L^2(\Omega)}^2 + \| \varphi_d \|_{L^2(\Omega)}^2 \big).$$
(5.7)

Then, substituting (5.5) and (5.7) into (5.4) yields

$$\frac{d}{dt} \int_{\Omega} F_{\varepsilon}(\varphi_{p},\varphi_{d}) + F_{1}(\varphi_{p},\varphi_{d}) + \frac{1}{2} \left(|\nabla\varphi_{p}|^{2} + |\nabla\varphi_{d}|^{2} \right) dx
+ \frac{M_{p}}{2} ||\nabla\mu_{p}||_{L^{2}(\Omega)}^{2} + \frac{M_{d}}{2} ||\nabla\mu_{d}||_{L^{2}(\Omega)}^{2} + \frac{1}{2} ||\boldsymbol{u}||_{L^{2}(\Omega)}^{2}
\leq C \left(1 + ||\varphi_{p}||_{H^{1}(\Omega)}^{2} + ||\varphi_{d}||_{H^{1}(\Omega)}^{2} + |(\mu_{p})_{\Omega}| + |(\mu_{d})_{\Omega}| \right).$$
(5.8)

The key point is now to derive uniform estimates on the mean values $|(\mu_p)_{\Omega}|$ and $|(\mu_d)_{\Omega}|$ in order to obtain useful a priori bounds from (5.8). To this aim, we test (5.1b) with $\varphi_p - (\varphi_p)_{\Omega}$ and (5.1d) with $\varphi_d - (\varphi_d)_{\Omega}$. Summing the resulting relations gives

$$\begin{aligned} |\nabla\varphi_p||^2_{L^2(\Omega)} + \|\nabla\varphi_d\|^2_{L^2(\Omega)} + \int_{\Omega} \nabla F_{\varepsilon}(\varphi_p,\varphi_d) \cdot (\varphi_p - (\varphi_p)_{\Omega},\varphi_d - (\varphi_d)_{\Omega})^{\top} dx \\ &\leq \int_{\Omega} (\mu_p - (\mu_p)_{\Omega})(\varphi_p - (\varphi_p)_{\Omega}) + (\mu_d - (\mu_d)_{\Omega})(\varphi_d - (\varphi_d)_{\Omega}) dx \\ &+ C \big(1 + \|\varphi_p\|_{L^2(\Omega)} + \|\varphi_d\|_{L^2(\Omega)} \big) \big(\|\nabla\varphi_p\|_{L^2(\Omega)} + \|\nabla\varphi_d\|_{L^2(\Omega)} \big). \end{aligned}$$
(5.9)

At this point we will use the fact that F_{ε} satisfies (3.2), and consider $s = \varphi_p$, $r = \varphi_d$, $S = (\varphi_p)_{\Omega}$, $R = (\varphi_d)_{\Omega}$. Then, we find that

$$c_* |\nabla F_{\varepsilon}(\varphi_p, \varphi_d) - \nabla F_{\varepsilon}((\varphi_p)_{\Omega}, (\varphi_d)_{\Omega})| \\ \leq (\nabla F_{\varepsilon}(\varphi_p, \varphi_d) - \nabla F_{\varepsilon}((\varphi_p)_{\Omega}, (\varphi_d)_{\Omega})) \cdot (\varphi_p - (\varphi_p)_{\Omega}, \varphi_d - (\varphi_d)_{\Omega})^{\top} + C_*.$$
(5.10)

Since $((\varphi_p)_{\Omega}, (\varphi_d)_{\Omega}) \in \Delta_0$ for all $t \in [0, T]$, we recall another property of the derivative of the Yosida approximation, namely

$$|\nabla F_{\varepsilon}(p,q)| \le |(\partial F_0)^{\circ}(p,q)| \quad \forall (p,q) \in \Delta,$$

where ∂ denotes here the subdifferential in the sense of convex analysis and $(\partial F_0)^{\circ}(p,q)$ is the element of minimum norm in the set $\partial F_0(p,q)$, that, at least in principle, could contain more than one element. Here, however, F_0 is assumed to be C^1 in Δ and, consequently, $|(\partial F_0)^{\circ}(p,q)| = |(\nabla F_0)(p,q)| < \infty$. Then, integrating (5.10) and rearranging leads to

$$\begin{aligned} c_* \|\nabla F_{\varepsilon}(\varphi_p, \varphi_d)\|_{L^1(\Omega)} &\leq c_* \|(\nabla F_0)((\varphi_p)_{\Omega}, (\varphi_d)_{\Omega})\|_{L^1(\Omega)} + C_* |\Omega| \\ &+ \int_{\Omega} \nabla F_{\varepsilon}(\varphi_p, \varphi_d) \cdot (\varphi_p - (\varphi_p)_{\Omega}, \varphi_d - (\varphi_d)_{\Omega})^\top dx \\ &+ C \|(\nabla F_0)((\varphi_p)_{\Omega}, (\varphi_d)_{\Omega})\|_{L^2(\Omega)} \big(\|\nabla \varphi_p\|_{L^2(\Omega)} + \|\nabla \varphi_d\|_{L^2(\Omega)} \big). \end{aligned}$$

Substituting this inequality into (5.9) then yields (cf. also [6])

$$\begin{aligned} \|\nabla\varphi_p\|_{L^2(\Omega)}^2 + \|\nabla\varphi_d\|_{L^2(\Omega)}^2 + \|\nabla F_{\varepsilon}(\varphi_p,\varphi_d)\|_{L^1(\Omega)} \\ &\leq \frac{M_p}{4} \|\nabla\mu_p\|_{L^2(\Omega)}^2 + \frac{M_d}{4} \|\nabla\mu_d\|_{L^2(\Omega)}^2 + C\left(1 + \|\varphi_p\|_{H^1(\Omega)}^2 + \|\varphi_d\|_{H^1(\Omega)}^2\right). \end{aligned}$$
(5.11)

Then, in light of (5.11), observe that, by testing (5.1b) and (5.1d) with ± 1 , we obtain

$$\begin{aligned} |(\mu_{p})_{\Omega}| + |(\mu_{d})_{\Omega}| &\leq C \left(1 + \|\varphi_{p}\|_{L^{2}(\Omega)} + \|\varphi_{d}\|_{L^{2}(\Omega)} \right) + \|\nabla F_{\varepsilon}(\varphi_{p},\varphi_{d})\|_{L^{1}(\Omega)} \\ &\leq \frac{M_{p}}{4} \|\nabla \mu_{p}\|_{L^{2}(\Omega)}^{2} + \frac{M_{d}}{4} \|\nabla \mu_{d}\|_{L^{2}(\Omega)}^{2} \\ &+ C \left(1 + \|\varphi_{p}\|_{H^{1}(\Omega)}^{2} + \|\varphi_{d}\|_{H^{1}(\Omega)}^{2} \right). \end{aligned}$$
(5.12)

Returning to (5.8) and substituting the estimate (5.12), we infer

$$\frac{d}{dt} \int_{\Omega} F_{\varepsilon}(\varphi_{p},\varphi_{d}) + F_{1}(\varphi_{p},\varphi_{d}) + \frac{1}{2} (|\nabla\varphi_{p}|^{2} + |\nabla\varphi_{d}|^{2}) dx
+ \frac{M_{p}}{4} ||\nabla\mu_{p}||_{L^{2}(\Omega)}^{2} + \frac{M_{d}}{4} ||\nabla\mu_{d}||_{L^{2}(\Omega)}^{2} + \frac{1}{2} ||\boldsymbol{u}||_{L^{2}(\Omega)}^{2}
\leq C (1 + ||\varphi_{p}||_{H^{1}(\Omega)}^{2} + ||\varphi_{d}||_{H^{1}(\Omega)}^{2}).$$
(5.13)

To (5.13) we now add the following inequality obtained from testing (5.1a) with φ_p and (5.1c) with φ_d and summing (cf. (3.31)):

$$\frac{1}{2}\frac{d}{dt}\left(\|\varphi_p\|_{L^2(\Omega)}^2 + \|\varphi_d\|_{L^2(\Omega)}^2\right) \le \frac{M_p}{8} \|\nabla\mu_p\|_{L^2(\Omega)}^2 + \frac{M_d}{8} \|\nabla\mu_d\|_{L^2(\Omega)}^2 + \frac{1}{4} \|\boldsymbol{u}\|_{L^2(\Omega)}^2 + C\left(1 + \|\varphi_p\|_{H^1(\Omega)}^2 + \|\varphi_d\|_{H^1(\Omega)}^2\right).$$

By definition of the Yosida approximation, we have

$$F_{\varepsilon}(s,r) \leq F_0(s,r) \quad \forall (s,r) \in \mathbb{R}^2.$$

Hence, recalling (2.9), we arrive at

$$\int_{\Omega} F_{\varepsilon}(\varphi_{p,0},\varphi_{d,0}) + F_1(\varphi_{p,0},\varphi_{d,0}) \, dx \le C.$$

Applying Gronwall's inequality in (5.13), we deduce

$$\begin{aligned} \|F_{\varepsilon}(\varphi_{p},\varphi_{d})\|_{L^{\infty}(0,T;L^{1}(\Omega))} + \|\varphi_{p}\|_{L^{\infty}(0,T;H^{1}(\Omega))} + \|\varphi_{d}\|_{L^{\infty}(0,T;H^{1}(\Omega))} \\ + \|\nabla\mu_{p}\|_{L^{2}(0,T;L^{2}(\Omega))} + \|\nabla\mu_{d}\|_{L^{2}(0,T;L^{2}(\Omega))} + \|\boldsymbol{u}\|_{L^{2}(0,T;L^{2}(\Omega))} \leq C. \end{aligned}$$
(5.14)

Thus, returning to (5.9), using the above estimate and performing some easy calculations, we infer

$$\|\nabla F_{\varepsilon}(\varphi_p, \varphi_d)\|_{L^1(\Omega)} \le C \left(1 + \|\nabla \mu_p\|_{L^2(\Omega)} + \|\nabla \mu_d\|_{L^2(\Omega)}\right), \tag{5.15}$$

whence $\nabla F_{\varepsilon}(\varphi_p, \varphi_d)$ is bounded in $L^2(0, T; L^1(\Omega))$. In turn, by the first line of (5.12) we find that $|(\mu_p)_{\Omega}|$ and $|(\mu_d)_{\Omega}|$ are bounded in $L^2(0, T)$. Hence, recalling (5.14) and using once more the Poincaré inequality, we get

$$\|\mu_p\|_{L^2(0,T;H^1(\Omega))} + \|\mu_d\|_{L^2(0,T;H^1(\Omega))} \le C.$$
(5.16)

Furthermore, recalling (5.6), thanks to (5.14) we now have

$$\|q\|_{L^2(0,T;H^1(\Omega))} \le C.$$
(5.17)

Next, we infer estimates on $F_{\varepsilon,p}$ by testing (5.1b) with $F_{\varepsilon,p}(\varphi_p,\varphi_d) - (F_{\varepsilon,p}(\varphi_p,\varphi_d))_{\Omega}$, leading to

$$\begin{split} \|F_{\varepsilon,p}(\varphi_p,\varphi_d) - (F_{\varepsilon,p}(\varphi_p,\varphi_d))_{\Omega}\|_{L^2(\Omega)}^2 \\ &+ \int_{\Omega} F_{\varepsilon,pp}(\varphi_p,\varphi_d) |\nabla \varphi_p|^2 + F_{\varepsilon,pd}(\varphi_p,\varphi_d) \nabla \varphi_p \cdot \nabla \varphi_d \, dx \\ &= \int_{\Omega} \left((\mu_p - (\mu_p)_{\Omega}) - F_{1,p}(\varphi_p,\varphi_d) \right) (F_{\varepsilon,p}(\varphi_p,\varphi_d) - (F_{\varepsilon,p}(\varphi_p,\varphi_d))_{\Omega}) \, dx, \end{split}$$

where

$$F_{\varepsilon,pp} = \frac{\partial^2 F_{\varepsilon}}{\partial \varphi_p^2}, \quad F_{\varepsilon,pd} = \frac{\partial^2 F_{\varepsilon}}{\partial \varphi_p \partial \varphi_d}.$$

Adding the similar identity obtained testing (5.1d) with $F_{\varepsilon,d}(\varphi_p,\varphi_d) - (F_{\varepsilon,d}(\varphi_p,\varphi_d))_{\Omega}$ and employing the Poincaré inequality together with the linear growth of ∇F_1 , it is not difficult to deduce

$$\begin{split} \|F_{\varepsilon,p}(\varphi_p,\varphi_d) - (F_{\varepsilon,p}(\varphi_p,\varphi_d))_{\Omega}\|_{L^2(\Omega)}^2 + \|F_{\varepsilon,d}(\varphi_p,\varphi_d) - (F_{\varepsilon,d}(\varphi_p,\varphi_d))_{\Omega}\|_{L^2(\Omega)}^2 \\ + \int_{\Omega} (\nabla\varphi_p,\nabla\varphi_d) \cdot D^2 F_{\varepsilon}(\varphi_p,\varphi_d) (\nabla\varphi_p,\nabla\varphi_d)^{\top} dx \\ &\leq C (1 + \|\nabla\mu_p\|_{L^2(\Omega)}^2 + \|\nabla\mu_d\|_{L^2(\Omega)}^2 + \|\varphi_p\|_{L^2(\Omega)}^2 + \|\varphi_d\|_{L^2(\Omega)}^2). \end{split}$$

Since F_{ε} is convex, the Hessian $D^2 F_{\varepsilon}$ is non-negative and consequently we can neglect the integral term on the left-hand side, leading to (cf. (5.14))

$$\begin{aligned} \|F_{\varepsilon,p}(\varphi_p,\varphi_d) - (F_{\varepsilon,p}(\varphi_p,\varphi_d))_{\Omega}\|_{L^2(0,T;L^2(\Omega))} \\ + \|F_{\varepsilon,d}(\varphi_p,\varphi_d) - (F_{\varepsilon,d}(\varphi_p,\varphi_d))_{\Omega}\|_{L^2(0,T;L^2(\Omega))} \le C. \end{aligned}$$
(5.18)

Upon recalling the boundedness of $\nabla F_{\varepsilon}(\varphi_p, \varphi_d)$ in $L^2(0, T; L^1(\Omega))$ resulting from (5.15), we deduce a control of the quantities $\|(F_{\varepsilon,p})_{\Omega}\|_{L^2(0,T)}$ and $\|(F_{\varepsilon,d})_{\Omega}\|_{L^2(0,T)}$. Hence, from (5.18) we eventually obtain

$$\|F_{\varepsilon,p}(\varphi_p,\varphi_d)\|_{L^2(0,T;L^2(\Omega))} + \|F_{\varepsilon,d}(\varphi_p,\varphi_d)\|_{L^2(0,T;L^2(\Omega))} \le C.$$
(5.19)

Viewing (5.1b) and (5.1d) as elliptic equations for φ_p and φ_d , respectively, with righthand sides bounded in $L^2(0,T;L^2)$ and no-flux boundary conditions, the elliptic regularity theory gives

$$\|\varphi_p\|_{L^2(0,T;H^2(\Omega))} + \|\varphi_d\|_{L^2(0,T;H^2(\Omega))} \le C.$$
(5.20)

Lastly, from inspection of (5.1a) and (5.1c), and thanks to the estimate (5.14) we have

$$\|\partial_t \varphi_p\|_{L^2(0,T;H^1(\Omega)')} + \|\partial_t \varphi_d\|_{L^2(0,T;H^1(\Omega)')} \le C.$$
(5.21)

5.2Compactness and passing to the limit

Thanks to the uniform estimates (5.14), (5.16), (5.17), (5.19), (5.20) and (5.21), by standard compactness arguments we infer the existence of functions $(\varphi_p, \mu_p, \varphi_d, \mu_d, q, u)$ and of a pair (η_p, η_d) such that

$$\varphi_i^{\varepsilon} \to \varphi_i \text{ weakly } \ast \text{ in } L^{\infty}(0,T;H^1(\Omega)) \cap L^2(0,T;H^2(\Omega)) \cap H^1(0,T;H^1(\Omega)'), \quad (5.22a)$$

$$\varphi_i^{\varepsilon} \to \varphi_i \text{ strongly in } C^0([0,T]; L^p(\Omega)) \cap L^2(0,T; W^{1,p}(\Omega)),$$

$$(5.22b)$$

 $\varphi_i^{\varepsilon} \to \varphi_i \text{ a.e. in } \Omega \times (0,T),$ $\mu_i^{\varepsilon} \to \mu_i \text{ weakly in } L^2(0,T;.)$ (5.22c)

$$\mu_i^{\varepsilon} \to \mu_i \text{ weakly in } L^2(0,T;H^1(\Omega)),$$
(5.22d)

$$\boldsymbol{u}^{\varepsilon} \to \boldsymbol{u} \text{ weakly in } L^2(0,T;L^2(\Omega)),$$
 (5.22e)

$$q^{\varepsilon} \to q \quad \text{weakly in } L^2(0,T;H^1(\Omega)),$$

$$(5.22f)$$

and

$$F_{\varepsilon,p}(\varphi_p,\varphi_d) \to \eta_p \text{ weakly in } L^2(0,T;L^2(\Omega)),$$
 (5.23a)

$$F_{\varepsilon,d}(\varphi_p,\varphi_d) \to \eta_d \text{ weakly in } L^2(0,T;L^2(\Omega)),$$
 (5.23b)

for any $p < \infty$ in two dimensions and any $p \in [1, 6)$ in three dimensions. Using a similar argument as in the proof of Theorem 3.4, by the a.e convergence of φ_p^{ε} to φ_p in $\Omega \times (0,T)$ and Egorov's theorem, we can show that $\{n^{\varepsilon}\}_{\varepsilon \in (0,1)}$ is a Cauchy family in $L^2(0,T; H^1(\Omega))$. Then, there also exists a function $n \in L^{\infty}(0,T; W^{2,r}(\Omega))$, for any $r < \infty$, with $0 \le n \le 1$ a.e. in $\Omega \times (0,T)$, such that

 $n^{\varepsilon} \to n$ strongly in $L^2(0,T; H^1(\Omega))$.

It now remains to pass to the limit $\varepsilon \to 0$ in (5.1). Actually, in view of the above convergence properties, the argument is very similar to that used before when we pass to the limit $\delta \to 0$. Hence, we just outline the differences which are mainly related to the terms depending on F_{ε} . Actually, combining (5.22b), (5.23a)-(5.23b) with the standard monotonicity argument in [2, Prop. 1.1, p. 42], we readily deduce that $(\varphi_p, \varphi_d) \in \Delta$ a.e. in $\Omega \times (0,T)$ and that $\eta_p = F_{0,p}(\varphi_p,\varphi_d), \ \eta_d = F_{0,d}(\varphi_p,\varphi_d)$. This in particular implies that the truncation operator $T(\cdot)$ disappears in the limit formulation of the problem; namely, we have $T(\varphi_p) = \varphi_p$ and $T(\varphi_d) = \varphi_d$ a.e. in Q.

Let us also point out that from the structural assumption (2.2) and from the derivation of (5.3) the limit functions φ_p and φ_d satisfy $((\varphi_p)_{\Omega}(t), (\varphi_d)_{\Omega}(t)) \in \Delta_0$ and

$$0 < c_1 \le (\varphi_p)_{\Omega}(t), (\varphi_d)_{\Omega}(t) \le c_2 < 1, \quad c_1 \le (\varphi_p + \varphi_d)_{\Omega}(t) \le c_2$$

for all $t \in [0,T]$. Hence, we have proved that the tuple $(\varphi_p, \mu_p, \varphi_d, \mu_d, \boldsymbol{u}, q, n)$ is a weak solution to system (1.6) in the sense of Definition 2.1. This concludes the proof of Theorem 2.1.

Acknowledgements

This research has been performed in the framework of the project Fondazione Cariplo-Regione Lombardia MEGAsTAR "Matematica d'Eccellenza in biologia ed ingegneria come acceleratore di una nuova strateGia per l'ATtRattività dell'ateneo pavese". The present paper also benefits from the support of the MIUR-PRIN Grant 2015PA5MP7 "Calculus of Variations" for GS, and of the GNAMPA (Gruppo Nazionale per l'Analisi Matematica, la Probabilità e le loro Applicazioni) of INdAM (Istituto Nazionale di Alta Matematica) for SF, ER, and GS. SF is "titolare di un Assegno di Ricerca dell'Istituto Nazionale di Alta Matematica".

References

- R.P. Aranjo and D.L.S. McElwain. A history of the study of solid tumor growth: the contribution of mathematical modelling, Bull. Math. Biol., 66 (2004), 1039–1091.
- [2] V. Barbu, "Nonlinear Semigroups and Differential Equations in Banach Spaces", Noordhoff, Leyden, 1976.
- [3] N. Bellomo, N.K. Li and P.K. Maini. On the foundations of cancer modelling: selected topics, speculations, and perspectives, Math. Models Methods Appl. Sci., 18 (2008), 593–646.
- [4] F. Boyer and S. Minjeaud. Hierarchy of consistent n-component Cahn-Hilliard systems, Math. Models Methods Appl. Sci., 24 (2014), 2885–2928.
- [5] D. Bresch, T. Colin, E. Grenier, B. Ribba and O. Saut. Computational modeling of solid tumor growth: the avascular stage, SIAM J. Sci. Comput., 32 (2010), 2321– 2344.
- [6] H. Brezis, "Opérateurs maximaux monotones et semi-groupes de contractions dans les espaces de Hilbert", North-Holland Math. Stud. 5, North-Holland, Amsterdam, 1973.
- [7] H.A. Bryne, T. Alarcón, M. Owen, S. Webb and P. Maini. Modeling aspects of cancer dynamics: a review, Philos. Trans. R. Soc. Lond. Ser. A Math. Phys. Eng. Sci., 364 (2006), 1563–1578.
- [8] A.C. Burton. Rate of growth of solid tumors as a problem of diffusion, Growth, 30 (1966), 157–176.
- [9] Y. Chen, S.M. Wise, V.B. Shenoy and J.S. Lowengrub. A stable scheme for a nonlinear, multiphase tumor growth model with an elastic membrane, Int. J. Numer. Meth. Biomed. Engng., **30** (2014), 726–754.
- [10] P. Colli and T. Fukao. Equation and dynamic boundary condition of Cahn-Hilliard type with singular potentials, Nonlinear Anal., 127 (2015), 413–433.
- [11] P. Colli, G. Gilardi and D. Hilhorst. On a Cahn-Hilliard type phase field model related to tumor growth, Discrete Contin. Dyn. Syst., 35 (2015), 2423–2442.
- [12] P. Colli, G. Gilardi, G. Marinoschi and E. Rocca. Sliding mode control for phase field system related to tumor growth, preprint arXiv:1706.03564 (2017), 1–22.
- [13] P. Colli, G. Gilardi, E. Rocca and J. Sprekels. Asymptotic analyses and error estimates for a Cahn-Hilliard type phase field system modelling tumor growth, Discrete Contin. Dyn. Syst. Ser. S., 10 (2017), 37–54.
- [14] P. Colli, G. Gilardi, E. Rocca and J. Sprekels. Optimal distributed control of a diffuse interface model of tumor growth, Nonlinearity, 30 (2017), 2518–2546.
- [15] P. Colli, G. Gilardi, E. Rocca and J. Sprekels. Vanishing viscosities and error estimate for a Cahn-Hilliard type phase field system related to tumor growth, Nonlinear Anal. Real World Appl., 26 (2015), 93–108.

- [16] P. Colli, P. Krejči, E. Rocca and J. Sprekels. Nonlinear evolution inclusions arising from phase change models, Czech Math. J., 57 (2007), 10067–1098.
- [17] M. Conti, S. Gatti and A. Miranville. Multi-component Cahn-Hilliard systems with dynamic boundary conditions, Nonlinear Anal. Real World Appl., 25 (2015), 137– 166.
- [18] V. Cristini, X. Li, J.S. Lowengrub and S.W. Wise. Nonlinear simulations of solid tumor growth using a mixture model: invasion and branching, J. Math. Biol., 58 (2009), 723–763.
- [19] V. Cristini and J. Lowengrub. Multiscale modeling of cancer. An Integrated Experimental and Mathematical Modeling Approach. Cambridge University Press, 2010.
- [20] M. Dai, E. Feireisl, E. Rocca, G. Schimperna and M. Schonbek. Analysis of a diffuse interface model for multispecies tumor growth, Nonlinearity, 30 (2017), 1639–1658.
- [21] A. Fasano, A. Bertuzzi and A. Gandolfi. Mathematical modeling of tumor growth and treatment. In A. Quarteroni, L. Formaggia and A. Veneziani, editors, Complex Systems in Biomedicine, pages 71-108. Springer, Milan, 2006.
- [22] H.B. Frieboes, F. Jin, Y.-L. Chuang, S.M. Wise, J.S. Lowengrub and V. Cristini. Three-dimensional multispecies nonlinear tumor growth - II: Tumor invasion and angiogenesis, J. Theor. Biol., 264 (2010), 1254–1278.
- [23] A. Friedman. Mathematical analysis and challenges arising from models of tumor growth, Math. Models Methods Appl. Sci., 17 (2007), 1751–1772.
- [24] S. Frigeri, M. Grasselli and E. Rocca. On a diffuse interface model of tumor growth, European J. Appl. Math., 26 (2015), 215–243.
- [25] S. Frigeri, K.F. Lam and E. Rocca. On a diffuse interface model for tumour growth with non-local interactions and degenerate mobilities, preprint arXiv:1703.03553 (2017), 1–28, "Solvability, Regularity, Optimal Control of Boundary Value Problems for PDEs", P. Colli, A. Favini, E. Rocca, G. Schimperna, J. Sprekels (eds.), Springer INdAM Series, Springer, Milan, 2017.
- [26] H. Garcke and K.F. Lam. Analysis of a Cahn-Hilliard system with non-zero Dirichlet conditions modeling tumor growth with chemotaxis, Discrete Contin. Dyn. Sys., 37 (2017), 4277–4308.
- [27] H. Garcke and K.F. Lam. Global weak solutions and asymptotic limits of a Cahn-Hilliard-Darcy system modelling tumour growth, AIMS Mathematics., 1 (2016), 318– 360.
- [28] H. Garcke and K.F. Lam. Well-posedness of a Cahn-Hilliard system modelling tumour growth with chemotaxis and active transport, European J. Appl. Math., 28 (2017), 284–316.
- [29] H. Garcke, K.F. Lam, R. Nürnberg and E. Sitka. A multiphase Cahn-Hilliard-Darcy model for tumour growth with necrosis, preprint arXiv:1701.06656 (2017), 1–43.
- [30] H. Garcke, K.F. Lam and E. Rocca. Optimal control of treatment time in a diffuse interface model for tumour growth, Appl. Math. Optim., (2017), DOI:10.1007/s00245-017-9414-4.

- [31] H. Garcke, K.F. Lam, E. Sitka and V. Styles. A Cahn-Hilliard-Darcy model for tumour growth with chemotaxis and active transport, Math. Models Methods Appl. Sci., 26 (2016), 1095–1148.
- [32] A. Giorgini, M. Grasselli and H. Wu. The Cahn-Hilliard-Hele-Shaw system with singular potential, Preprint HAL, Id: hal-01543386 (2017), submitted.
- [33] H.P. Greenspan. Models for the growth of a solid tumor by diffusion, Studies in Applied Mathematics, **51** (1972), 317–340.
- [34] A. Hawkins-Daarud, K.G. van der Zee and J.T. Oden. Numerical simulation of a thermodynamically consistent four-species tumor growth model, Int. J. Numer. Methods Biomed. Eng., 28 (2012), 3–24.
- [35] J. Jiang, H. Wu and S. Zheng. Well-posedness and long-time behavior of a nonautonomous Cahn-Hilliard-Darcy system with mass source modeling tumor growth, J. Differential Equations, 259 (2015), 3032–3077.
- [36] Y. Li, J.-I. Choi and J. Kim. Multi-component Cahn-Hilliard system with different boundary conditions in complex domains, J. Comput. Phys., 323 (2016), 1–16.
- [37] J.S. Lowengrub, E. Titi and K. Zhao. Analysis of a mixture model of tumor growth, European J. Appl. Math., 24 (2013), 691–734.
- [38] S. Melchionna and E. Rocca. Varifold solutions of a sharp interface limit of a diffuse interface model for tumor growth, preprint arXiv:1610.04478 (2016), 1–21, Interfaces and Free Boundaries, to appear (2017).
- [39] A. Miranville and S. Zelik. The Cahn-Hilliard equation with singular potentials and dynamic boundary conditions, Discrete Contin. Dyn. Syst., 28 (2010), 275–310.
- [40] J.T. Oden, A. Hawkins and S. Prudhomme. General diffuse-interface theories and an approach to predictive tumor growth modeling, Math. Models Methods Appl. Sci., 58 (2010), 723–763.
- [41] E. Rocca and R. Scala. A rigorous sharp interface limit of a diffuse interface model related to tumor growth, J. Nonlinear Sci, 27 (2017), 847–872.
- [42] G. Sciumè, S. Shelton, W.G. Gray, C.T. Miller, F. Hussain, M. Ferrari, P. Decuzzi and B.A. Schrefler. A multiphase model for three-dimensional tumor growth, New Journal of Physics, 15 (2013), 015005.
- [43] J. Simon. Compact sets in the space $L^p(0,T;B)$, Ann. Mat. Pura Appl., **146** (1986), 65–96.
- [44] S.M. Wise, J.S. Lowengrub, H.B. Frieboes and V. Cristini. Three-dimensional multispecies nonlinear tumor growth - I: model and numerical method, J. Theor. Biol., 253 (2008), 524–543.
- [45] H. Youssefpour, X. Li, A.D. Lander and J.S. Lowengrub. Multispecies model of cell lineages and feedback control in solid tumors, J. Theor. Biol., 304 (2012), 39–59.