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1	An Asymptotic Homogenization Approach to the
2	Microstructural Evolution of Heterogeneous Media
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14 Abstract

In the present work, we apply the asymptotic homogenization technique to the equations describing the dynamics of a heterogeneous material with evolving micro-structure, thereby obtaining a set of upscaled, effective equations. We consider the case in which the heterogeneous body comprises two hyperelastic materials and we assume that the evolution of their micro-structure occurs through the development of plastic-like distortions, the latter ones being accounted for by means of the Bilby-Kröner-Lee (BKL) decomposition. The asymptotic homogenization approach is applied simultaneously to the linear momentum balance law of the body and to the evolution law for the plastic-like distortions. Such evolution law models a stress-driven production of inelastic distortions, and stems from phenomenological observations done on cellular aggregates. The whole study is also framed within the limit of small elastic distortions, and provides a robust framework that can be readily generalized to growth and remodeling of nonlinear composites. Finally, we complete our theoretical model by performing numerical simulations.

- ¹⁵ Keywords: Asymptotic homogenization, heterogeneous media, remodeling,
- ¹⁶ BKL decomposition, two-scale plasticity, nonlinear composites

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17 1. Introduction

The study of material growth, remodeling and aging is of great importance in Biomechanics, specially when the tissue, in which these processes occur, features a very complex structure, with different scales of observation and various constituents.

In the literature, the study of heterogeneous materials follows several 22 approaches. In this work we focus on the multi-scale asymptotic homoge-23 nization technique [4, 5, 8, 14, 77], which exploits the information available 24 at the smallest scale characterizing the considered medium or phenomenon to 25 obtain an effective description of the medium or phenomenon itself valid at 26 its largest scale. This is achieved by expanding in asymptotic series the equa-27 tions constituting the mathematical model formulated at the lowest scale. As 28 a result, the coefficients of the effective governing equations encode the infor-20 mation on the other hierarchical levels, as they are to be computed solving 30 microstructural problems at the smaller scales. The multi-scale asymptotic 31 homogenization approach has been successfully applied to investigate var-32 ious physical systems due to its potentiality in decreasing the complexity 33 of the problem at hand. Biomechanical applications of asymptotic homoge-34 nization may be found mainly in nanomedicine [81], biomaterials modeling, 35 such as the bone [58, 65], tissue engineering [24], poroelasticity [63], and ac-36 tive elastomers [64]. Most of the literature concerning applications of the 37 asymptotic homogenization technique focuses on linearized governing equa-38 tions, as in this case it is possible to obtain, under a number of simplifying 39 assumptions, a full decoupling between scales, which leads to a dramatic re-40 duction in the computational complexity, as also noted for example in [64]. 41 In fact, homogenization in nonlinear mechanics is usually tackled via average 42 field approaches based on representative volume elements or Eshelby-based 43 techniques (see e.g. [41] for a comparison between the latter and asymp-44 totic homogenization), as done for example in [11]. These homogenization 45 approaches are typically well-suited when seeking for suitable bounds for the 46 coefficients of the model, such as the elastic moduli, while asymptotic ho-47 mogenization can provide a precise characterization of the coefficients under 48 appropriate regularity assumptions (namely, *local periodicity*). 49

However, to the best of our knowledge and understanding, there exists only a few examples, e.g. [15, 68, 74, 75], dealing with the asymptotic homogenization in the case of media undergoing large deformations. In [68], the static microstructural effects of periodic hyperelastic composites at finite

strain are investigated. In [74], the interactions between large deforming solid 54 and fluid media at the microscopic level are described by using the two-scale 55 homogenization technique and the updated Lagrangian formulation. In [15], 56 the effective equations describing the flow, elastic deformation and transport 57 in an active poroelastic medium were obtained. Therein, the authors consid-58 ered the spatial homogenization of a coupled transport and fluid-structure 59 interaction model, incorporating details of the microscopic system and ad-60 mitting finite growth and deformation at the pore scale. Some works can be 61 also found dealing with homogenization in the case of elastic perfectly plastic 62 constituents [79, 83]. 63

Here we embrace the asymptotic homogenization approach and consider 64 a heterogeneous body composed of two hyperelastic solid constituents sub-65 jected to the evolution of their internal structure. We refer to this phe-66 nomenon as to material remodeling and we interpret it with the production 67 of plastic-like distortions. The wording "material remodeling" is used as a 68 synonym of "evolution of the internal structure" of a tissue, and is intended in 69 the sense of [16], who states that "biological systems can adapt their structure 70 [...] to accommodate a changed mechanical load environment". In this case, 71 always in the terminology of [16] and [80], one speaks of *epigenetic* adap-72 tation (or material remodeling). In the framework of the manuscript, such 73 adaptation is assumed to occur through plastic-like distortions that represent 74 processes like the redistribution of the adhesion bonds among the tissue cells. 75

It is worth to recall in which sense the concept of "plastic distortions". 76 conceived in the context of the Theory of Plasticity (cf. e.g. [50, 55]). 77 and originally referred to non-living materials such as metals or soils, can 78 be imported to describe the structural evolution of biological tissues. To 79 this end, it is important to emphasize that the wording "plastic distortions" 80 is understood as the result of a complex of transformations that conducts 81 to the reorganization of the internal structure of a material, and that -82 as anticipated in the Introduction— such reorganization is referred to as 83 "remodeling" in the biomechanical context. 84

The ways in which the structural tranformations may take place in a given material depend on the structural properties of the material itself. For this reason, the plasticity in metals is markedly different from that occurring in amorphic materials. In the case of metals, indeed, for which the internal structure is granular and characterized by the arrangement of the atomic lattice within each grain, plastic distortions are the *macroscopic* manifestation of the formation and evolution of lattice defects. As reported in [55], such

defects can be due, for example, to edge dislocations, wedge disclinations, 92 missing atoms at some lattice sites, or to the presence of atoms in the lat-93 tice interstices. To describe how the defects evolve, thereby giving rise to the 94 plastic distortions, one should compare the real lattice at the current instant 95 of time with an ideal lattice, and decompose the overall deformation (i.e., 96 shape change and structural transformation) into an elastic and an inelastic 97 contribution [55]. The elastic contribution describes the part of deformation 98 that is recoverable by completely relaxing mechanical stress, whereas the in-99 elastic contribution represents the structural variation, which, in general, is 100 of irreversible nature. 101

¹⁰² Clearly, metals have structural features markedly different from those of ¹⁰³ living matter. Still, some of the fundamental mechanisms that trigger the ¹⁰⁴ reorganization of their internal structure can be adapted to describe the ¹⁰⁵ remodeling of biological tissues.

For instance, in the case of bones, plastic-like phenomena are due to 106 the formation of microcracks that, in turn, favors the gliding of the material 107 along the direction of the opening of the cracks [17, 86]. Lastly, as anticipated 108 above, in the case of biological tissues such as cellular aggregates, the phe-109 nomenon analogous to the generation of dislocations is the rearrangement of 110 the adhesion bonds among the cells or the reorganization of the extracellular 111 matrix due to the reorientation of the collagen fibers or their deposition and 112 resorption, as is the case for blood vessels [48]. Also in all these situations, 113 the comparison of the real configuration of the tissue with an "ideal" one, 114 taken as reference, permits the separation of the overall deformation into an 115 elastic part and a structure-related, "plastic-like" part. 116

Here, taking inspiration from the theory of finite Elastoplasticity [55, 78, 117 34, we describe the plastic-like distortions by invoking the Bilby-Kröner-Lee 118 (BKL) decomposition of the deformation gradient tensor, and rephrasing it in 119 a scale-dependent fashion. We remark that, at each of the medium's charac-120 teristic scales, a tensor of plastic distortions is introduced, which accounts for 121 the fact that the structural variations of the medium cannot be expressed, in 122 general, in terms of compatible deformations. Our study is conducted within 123 a purely mechanical framework and under the assumption of negligible iner-124 tial forces. These hypotheses imply that the model equations reduce to a set 125 comprising a scale-dependent, quasi-static law of balance of linear momen-126 tum and an evolution law for the tensor of plastic-like distortions. The latter 127 one is assumed to obey a phenomenological flow rule driven by stress. 128

¹²⁹ The manuscript is organized as follows. In Section 2, we introduce the

fundamental notions related to the separation of scales, kinematics, and the 130 Bilby-Kröner-Lee decomposition for the heterogeneous material. Therein, 131 the kinematics of the considered medium is discussed, which has to account 132 for the different length-scales characterizing the heterogeneities and results 133 into the definition of a scale-dependent deformation gradient tensor. In Sec-134 tion 3, the problem to be solved is formulated, and in Section 4, the two-135 scales asymptotic homogenization technique is applied to obtain the local 136 and the homogenized sub-problems. In Section 5, we prescribe a constitutive 137 equation for the response of the material and, independently, an evolution 138 equation for the tensor of plastic-like distortions. In that respect, the local 139 and homogenized problems derived in Section 4 are formulated by consid-140 ering the De Saint-Venant strain energy density and we demonstrate the 141 relationship between our new model and the classical ones. In Section 6 we 142 outline a computational scheme to solve the resulting up-scaled model and, 143 in Section 7, we address the numerical results of our simulations. Finally, 144 some concluding remarks on the ongoing work, along with suggestions for 145 future research, are summarized in Section 8. We highlight the novelty of 146 our approach, and we explain how it may contribute to the understanding of 147 the mechanics of heterogeneous media with evolving micro-structure. 148

¹⁴⁹ 2. Theoretical background

150 2.1. Separation of scales

The homogenization of a highly heterogeneous medium is only possible when the characteristic length of the the local structure (ℓ_0) and the characteristic length of the material, or of the phenomenon, of interest (L_0) are well separated. This condition of separation of scales can be expressed as

$$\varepsilon_0 := \frac{\ell_0}{L_0} \ll 1. \tag{1}$$

There may exist more than two coexisting scales and, if they are well separated from each other, a homogenization approach is possible. In this case, we then move from the smallest scale to the largest one by homogenization [1, 8, 51, 82, 69].

¹⁵⁹ Condition (1) is taken as a base assumption for all homogenization pro-¹⁶⁰ cesses. The two characteristic length scales ℓ_0 and L_0 introduce two di-¹⁶¹ mensionless spatial variables in the reference configuration, $\tilde{Y} = X/\ell_0$ and ¹⁶² $\tilde{X} = X/L_0$, where X is said to be the *physical spatial variable*, whereas \tilde{Y} and \tilde{X} represent the microscopic and the macroscopic non-dimensional spatial variables, respectively. By using (1), \tilde{Y} and \tilde{X} can be related through the expression

$$\tilde{Y} = \varepsilon_0^{-1} \tilde{X}.$$
(2)

Given a field Φ defined over the region of interest of the heterogeneous medium, the separation of scales allows to rephrase the space dependence of Φ as $\Phi(X) = \check{\Phi}(\tilde{X}(X), \tilde{Y}(X))$, and the spatial derivative of Φ takes thus the form

$$\operatorname{Grad}_{X} \Phi = L_{0}^{-1} \left(\operatorname{Grad}_{\tilde{X}} \check{\Phi} + \varepsilon_{0}^{-1} \operatorname{Grad}_{\tilde{Y}} \check{\Phi} \right).$$
(3)

By following this approach, all equations should be written in non-dimensional 170 form. In the literature, the switch to the auxiliary variables \tilde{X} and \tilde{Y} is often 171 omitted. However, as shown for example in [4], both paths are equivalent, 172 provided that the dimensional formulation of the problem consistently ac-173 counts for any asymptotic behavior of the involved fields and parameters 174 (see e.g. [62] and the discussion therein concerning problems where such a 175 behavior is actually deduced via a non-dimensional analysis). By exploiting 176 this result, in what follows, our analysis is carried out directly in a system of 177 physical variables X and Y. Moreover, by adopting the approach usually fol-178 lowed in asymptotic multiscale analysis, we assume that each field and each 179 material property characterizing the considered medium are functions of both 180 X and Y, with $Y = \varepsilon_0^{-1} X$. Roughly speaking, the dependence on X captures 181 the behavior of a given physical quantity over the largest length-scale, while 182 the dependence on Y captures the behavior over the smallest one. We express 183 this property by introducing the notation $\Phi^{\varepsilon}(X) = \Phi(X, \varepsilon_0^{-1}X) = \Phi(X, Y)$ 184 [66]. Moreover, for a fixed X, we assume that $\Phi(X,Y)$ is periodic with 185 respect to Y. 186

In the classical theory of two-scale asymptotic homogenization [5, 8, 14], 187 the small scaling dimensionless parameter ε_0 is constant. However, in the 188 case of a composite material subjected to deformation and change of internal 189 structure (as is the case, for instance, when plastic-like distortions occur), 190 the characteristic macroscopic and microscopic lengths, which refer to the 191 body and to its heterogeneities, respectively, depend on X and t, and should 192 thus be denoted by $\ell(X,t)$ and L(X,t). Therefore, the corresponding scaling 193 parameter, obtained as the ratio $\varepsilon(X,t) = \ell(X,t)/L(X,t)$, is also a func-194 tion of X and t, which need not be equal to ε_0 in general. This variability 195

notwithstanding, if $\varepsilon(X, t)$ is bounded from above for all X and for all t, and if the upper bound is much smaller than unity, we can indicate such upper bound with ε , and use this constant as a scaling parameter for our asymptotic analysis.

200 2.2. Kinematics

Let us denote by $\mathcal{B}^{\varepsilon}$ a continuum body with periodic microstructure, and 201 by \mathcal{S} the three-dimensional Euclidean space. Furthermore, we denote by 202 $\mathcal{B}_0^{\varepsilon}$ the reference, unloaded configuration of $\mathcal{B}^{\varepsilon}$, in which the body's periodic 203 micro-structure is reproduced. Now, let us assume that $\chi^{\varepsilon} : \mathcal{B}_{0}^{\varepsilon} \times \mathcal{T} \to \mathcal{S}$ 204 describes the motion of the heterogeneous body, where $\mathcal{T} = [t_0, t_f]$ is an 205 interval of time. Then, the region occupied by the body at time $t \in \mathcal{T}$ 206 is $\mathcal{B}_t^{\varepsilon} := \chi^{\varepsilon}(\mathcal{B}_0^{\varepsilon}, t) \subset \mathcal{S}$ and is said to be its current configuration. Each 207 point $x \in \mathcal{B}_t^{\varepsilon}$ is such that $x = \chi^{\varepsilon}(X, t)$, with $X \in \mathcal{B}_0^{\varepsilon}$ being the point's 208 reference placement. The deformation from $\mathcal{B}_0^{\varepsilon}$ to $\mathcal{B}_t^{\varepsilon}$ is characterized by the 209 deformation gradient, $\mathbf{F}^{\varepsilon}(X,t)$, which is defined as $\mathbf{F}^{\varepsilon}(X,t) = T\chi^{\varepsilon}(X,t)$ 210 [53], with $T\chi^{\varepsilon}$ being the tangent map of the motion χ^{ε} , defined from the 211 tangent space $T_X \mathcal{B}^{\varepsilon}_{\varepsilon}$ into $T_x \mathcal{S}$. In the sequel, however, since our focus is on 212 Homogenization Theory, we find it convenient to use the less formal definition 213

$$\boldsymbol{F}^{\varepsilon} = \boldsymbol{I} + \operatorname{Grad} \boldsymbol{u}^{\varepsilon}, \tag{4}$$

where I is the second-order identity tensor and $\operatorname{Grad} u^{\varepsilon}$ denotes the gradient 214 operator of the displacement $\boldsymbol{u}^{\varepsilon}$. The condition $J^{\varepsilon} = \det \boldsymbol{F}^{\varepsilon} > 0$ must be 215 satisfied in order for χ^{ε} to be admissible. The symmetric, positive definite, 216 second-order tensor $C^{\varepsilon} = (F^{\varepsilon})^T F^{\varepsilon}$ is the right Cauchy-Green deformation 217 tensor induced by F^{ε} . For our purposes, we partition $\mathcal{B}_0^{\varepsilon}$ into two sub-218 domains \mathcal{B}_0^1 and \mathcal{B}_0^2 , such that $\bar{\mathcal{B}}_0^1 \cup \bar{\mathcal{B}}_0^2 = \bar{\mathcal{B}}_0^{\varepsilon}$ and $\bar{\mathcal{B}}_0^1 \cap \mathcal{B}_0^2 = \mathcal{B}_0^1 \cap \bar{\mathcal{B}}_0^2 = \emptyset$, 219 where the bar over a set denotes its closure. We let Γ_0^{ε} stand for the interface 220 between \mathcal{B}_0^1 and \mathcal{B}_0^2 . Particularly, \mathcal{B}_0^1 denotes the matrix of $\mathcal{B}^{\varepsilon}$ (also referred 221 to as host phase) and \mathcal{B}_0^2 a collection of N disjoint inclusions. The periodic 222 cell in the reference configuration is denoted by \mathcal{Y}_0 . The portion of matrix 223 contained in \mathcal{Y}_0 is indicated by \mathcal{Y}_0^1 , while \mathcal{Y}_0^2 is the inclusion in \mathcal{Y}_0 . In each 224 cell, \mathcal{Y}_0^1 and \mathcal{Y}_0^2 are such that $\overline{\mathcal{Y}}_0^1 \cup \overline{\mathcal{Y}}_0^2 = \overline{\mathcal{Y}}_0$ and $\overline{\mathcal{Y}}_0^1 \cap \mathcal{Y}_0^2 = \mathcal{Y}_0^1 \cap \overline{\mathcal{Y}}_0^2 = \emptyset$. The 225 symbol Γ_0 indicates the interface between \mathcal{Y}_0^1 and \mathcal{Y}_0^2 . In the present work, we 226 assume that the periodicity of the body's micro-structure is preserved even 227 though the body evolves by both changing its shape and varying its internal 228 structure. In general, however, this is not the case. Clearly, our hypothesis is 229

unrealistic in several circumstances, but it might be helpful to describe those 230 situations in which the breaking of the material symmetries occurs at a scale 231 different from those of interest, as is the case, for instance, when the plastic 232 distortions occur in a tissue with evolving material properties [49], that are 233 not directly related to the change of the tissue's micro-geometry. On the 234 other hand, for nonperiodic media, the macro model is still valid when one 235 assumes local boundedness. In that case, the coefficients are simply to be 236 retrieved experimentally, as the "cell" problem is no longer to be computed 237 on the cell but on the whole micro domain, which would be more complex 238 than the original problem. 239

Moreover, we define $\chi_1^{\varepsilon} := \chi^{\varepsilon}|_{\mathcal{B}_0^1} : \mathcal{B}_0^1 \times \mathcal{T} \to \mathcal{S}$ such that $\mathcal{B}_t^1 := \chi_1^{\varepsilon}(\mathcal{B}_0^1, t)$ 240 denotes the host phase at the current configuration and $\chi_2^{\varepsilon} := \chi^{\varepsilon}|_{\mathcal{B}^2_0} : \mathcal{B}^2_0 \times$ 241 $\mathcal{T} \to \mathcal{S}$, with $\mathcal{B}_t^2 := \chi_2^{\varepsilon}(\mathcal{B}_0^2, t)$ denoting the inclusions. Specifically, we enforce 242 the condition $\bar{\mathcal{B}}_t^1 \cup \bar{\mathcal{B}}_t^2 = \bar{\mathcal{B}}_t^{\varepsilon}$, with $\bar{\mathcal{B}}_t^1 \cap \bar{\mathcal{B}}_t^2 = \mathcal{B}_t^1 \cap \bar{\mathcal{B}}_t^2 = \emptyset$, and denote by Γ_t^{ε} the 243 interface between \mathcal{B}_t^1 and \mathcal{B}_t^2 . In addition, we let \mathcal{Y}_t indicate the periodic cell 244 in the current configuration, with $\bar{\mathcal{Y}}_t^1 \cup \bar{\mathcal{Y}}_t^2 = \bar{\mathcal{Y}}_t$, $\bar{\mathcal{Y}}_t^1 \cap \mathcal{Y}_t^2 = \mathcal{Y}_t^1 \cap \bar{\mathcal{Y}}_t^2 = \emptyset$, and with Γ_t being the interface between \mathcal{Y}_t^1 and \mathcal{Y}_t^2 (see Fig. 1). We emphasize 245 246 that \mathcal{Y}_t^1 is the portion of matrix and \mathcal{Y}_t^2 is the inclusion in \mathcal{Y}_t . We note that 247 inside a single cell it can be present also a collection of inclusions and, in 248 such a case, we should consider multiple interface conditions [60]. 249

250 2.3. Multiplicative decomposition

When the body $\mathcal{B}^{\varepsilon}$ is subjected to a system of external loads, the change 251 of its shape could be accompanied by a rearrangement of its intrinsic struc-252 ture. This process is generally inelastic and may not be described just in 253 terms of deformation. Moreover, when mechanical agencies are removed, the 254 body is generally unable to recover the unloaded configuration $\mathcal{B}_{0}^{\varepsilon}$, and may 255 occupy a configuration characterized by the presence of residual stresses and 256 strains. To bring the body into a fully relaxed state, an ideal tearing process 257 has to be introduced [55]. More specifically, for each material point $X \in \mathcal{B}^{\varepsilon}$, 258 we individuate a small neighborhood of X, referred to as *body element*, we 259 ideally cut it out from the body, and we let it relax until it reaches a stress-260 free state. Such state is the ground state of the relaxed body element and 261 is called *natural state*. This concept, originally used in the theory of elasto-262 plasticity (see [50, 55]), has been used in the biomechanical context by various 263 authors like, for instance, [23, 76, 30, 26, 27, 42, 44, 18, 55, 34, 19]. Before 264 going further with the use of the BKL decomposition, we mention that, in 265 the literature, there exist other approaches to the issue of residual stresses in 266

biological tissues, which call neither for the multiplicative decomposition of 267 the deformation gradient tensor, nor for the introduction of an "intermediate, 268 relaxed configuration". One recent publication adhering to this philosophy 269 is for example [13], in which the authors warn that the intermediate config-270 uration may "not exist in physical reality and must be postulated a priori". 271 Although we are aware of the fact that a framework based on the BKL-272 decomposition may lead in some cases to assume unrealistic results —as any 273 other framework would do—, we prefer here to adhere to the BKL approach 274 for consistency with previous works of ours. 275

By performing the ideal process described above for all the body points, a 276 collection of relaxed body pieces is obtained, in which each piece finds itself 277 in its natural state. We denote such collection by $\mathcal{B}_{\nu}^{\varepsilon}$. In the language of 278 continuum mechanics, these physical considerations lead to the BKL decom-279 position [55, 34]. Although summarizing these theoretical results is useful for 280 sake of completeness, the consequences of the BKL decomposition are well-281 known, as it is one the pillars of Elastoplasticity. For this reason, we do not 282 fuss over its theoretical justification, and we highlight, rather, the fact that 283 one of the purposes of this work is to investigate the use of a scale-dependent 284 BKL decomposition. In detail, by referring to Figure 1, we invoke a multi-285 plicative decomposition of the deformation gradient F^{ε} that is parameterized 286 by the scaling ratio ε , i.e., 287

$$\boldsymbol{F}^{\varepsilon} = \boldsymbol{F}_{\mathrm{e}}^{\varepsilon} \boldsymbol{F}_{\mathrm{p}}^{\varepsilon},\tag{5}$$

where the tensors $\mathbf{F}_{e}^{\varepsilon}$ and $\mathbf{F}_{p}^{\varepsilon}$ describe, respectively, the elastic and the inelastic distortions contributing to \mathbf{F}^{ε} Along with (5), we also define the determinants $J_{e}^{\varepsilon} = \det(F_{e}^{\varepsilon})$ and $J_{p}^{\varepsilon} = \det(F_{p}^{\varepsilon})$, which are both strictly positive. Consistently with the notation introduced above, it holds true that $\mathbf{F}_{e}^{\varepsilon}(X) = \mathbf{F}_{e}(X,Y), \ \mathbf{F}_{p}^{\varepsilon}(X) = \mathbf{F}_{p}(X,Y), \ \text{and} \ \mathbf{F}^{\varepsilon}(X) = \mathbf{F}(X,Y)$ as well as $J_{e}^{\varepsilon}(X) = J_{e}(X,Y)$ and $J_{p}^{\varepsilon}(X) = J_{p}(X,Y).$

In this work, we focus on remodeling, i.e., plastic-like distortions that occur to modify the internal structure of $\mathcal{B}^{\varepsilon}$. Although this phenomenon is not visible, it could lead to the alteration of the mechanical properties of $\mathcal{B}^{\varepsilon}$.

²⁹⁷ 3. Formulation of the problem

We consider a composite material comprising two solid constituents, whose point-wise constitutive response is hyperelastic. Therefore, to model its mechanical behavior, we introduce the scale-dependent strain energy function,

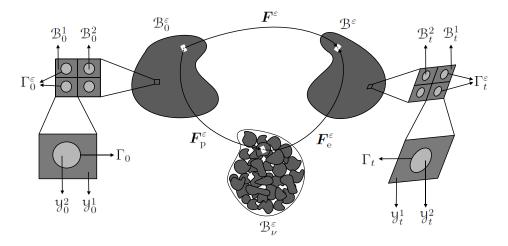


Figure 1: Schematic of a composite material with periodic internal micro-structure and subjected to inelastic remodeling distortions. From left to right: Magnification of an excerpt of material and description of its nested, periodic micro-structure. Change of shape of the body from the reference to the current configuration, and definition of the conglomerate of relaxed body pieces, each in its natural state. Magnification of an excerpt of material, taken from the body's current configuration, and description of its deformed, and remodeled, micro-structure.

301 defined per unit volume of the natural state,

$$\check{\psi}_{\nu}(X,t) = \psi_{\nu}^{\varepsilon}(\boldsymbol{F}_{e}^{\varepsilon}(X,t), i^{\varepsilon}(X,t)) = \psi_{\nu}(\boldsymbol{F}_{e}(X,Y,t), i(X,Y,t)), \qquad (6)$$

where *i* is defined by the expression i(X, Y, t) = (X, Y), i.e., *i* extracts the spatial pair (X, Y) from the triplet (X, Y, t). From (6) we can derive the first Piola-Kirchoff stress tensor,

$$\boldsymbol{T}^{\varepsilon} = J_{\mathrm{p}}^{\varepsilon} \frac{\partial \psi_{\nu}^{\varepsilon}}{\partial \boldsymbol{F}_{\mathrm{e}}^{\varepsilon}} \left(\boldsymbol{F}_{\mathrm{p}}^{\varepsilon} \right)^{-T}, \qquad (7)$$

where $J_{\rm p}^{\varepsilon} = \det F_{\rm p}^{\varepsilon}$. In particular, if we neglect body forces and inertial terms, the balance of linear momentum reads,

$$\begin{cases} \operatorname{Div} \boldsymbol{T}^{\varepsilon} = \boldsymbol{0}, & \operatorname{in} \ \mathcal{B}_{0}^{\varepsilon} \setminus \Gamma_{0}^{\varepsilon} \times \mathcal{T}, \\ \boldsymbol{T}^{\varepsilon} \cdot \boldsymbol{N} = \bar{\boldsymbol{T}}, & \operatorname{on} \ \partial_{T} \mathcal{B}_{0}^{\varepsilon} \times \mathcal{T}, \\ \boldsymbol{u}^{\varepsilon} = \bar{\boldsymbol{u}}, & \operatorname{on} \ \partial_{u} \mathcal{B}_{0}^{\varepsilon} \times \mathcal{T}, \end{cases}$$
(8)

where \bar{T} and \bar{u} are, respectively, the prescribed traction and displacement on the boundary $\partial \mathcal{B}_0^{\varepsilon} = \partial_T \mathcal{B}_0^{\varepsilon} \cup \partial_u \mathcal{B}_0^{\varepsilon}$ with $\overline{\partial_T \mathcal{B}_0^{\varepsilon}} \cap \partial_u \mathcal{B}_0^{\varepsilon} = \partial_T \mathcal{B}_0^{\varepsilon} \cap \overline{\partial_u \mathcal{B}_0^{\varepsilon}} = \emptyset$ and N is the outward unit vector normal to the surface $\partial \mathcal{B}_0^{\varepsilon}$. Continuity conditions for displacement and traction are imposed,

$$\llbracket \boldsymbol{u}^{\varepsilon} \rrbracket = \boldsymbol{0} \quad \text{and} \quad \llbracket \boldsymbol{T}^{\varepsilon} \cdot \boldsymbol{N}_{\mathcal{Y}} \rrbracket = \boldsymbol{0}, \quad \text{on } \Gamma_0 \times \mathcal{T}, \tag{9}$$

where $[\bullet]$ denotes the jump across the interface between the two constituents 311 and $N_{\mathcal{Y}}$ defines the unit outward normal to Γ_0 . Moreover, problem (8) 312 must be supplemented with an appropriate evolution law for $F_{\rm p}^{\varepsilon}$. It is worth 313 mentioning that the homogenization process can be performed regardless of 314 the particular choice of *external* boundary conditions (Dirichlet-Neumann 315 in this case). This means that the formulation presented in this work is 316 potentially applicable also to other external boundary conditions, such as 317 e.g. those of Robin-type. This is due to the fact that, as pointed out in [69], 318 also in the present study the homogenization is applied in regions sufficiently 319 far away from the outer boundary of the considered medium. For problems 320 in which it is necessary to homogenize also close to the outer heterogeneous 321 boundaries, we refer to [8, 57, 46]. 322

Remark 1. In the present work, we impose conditions (9) for displacements 323 and tractions just to exemplify the homogenization technique applied to het-324 erogeneous media with evolving microstructure. In other words, we assume 325 that the contact interface between the constituents is ideal. This means that 326 the displacements are congruent, and thus continuous, and that linear mo-327 mentum is conserved across the interface, which in our context implies the 328 continuity of the tractions. However, the hypothesis of the ideal interface can 329 be relaxed in some biological situations. For instance, in cancerous tissues, 330 there exist cross-links between normal and malignant cells, whose density and 331 strength determine a spring constant that relates the normal stresses on each 332 cell surface, thereby making it non-ideal [47, 37]. Another example of non-333 ideal interface is the periodontal ligament, which represents the thin layer 334 between the cementum of the tooth to the adjacent alveolar bone [28]. In the 335 context of composite materials, when non-ideal interfaces are accounted for, 336 the interface conditions are suitably reformulated [38, 39, 7, 6]. In particular, 337 the asymptotic homogenization technique has been applied for linear elastic 338 periodic fiber reinforced composites with imperfect contact between matrix and 339 fibers (see e.g. [36]). 340

³⁴¹ 4. Asymptotic homogenization of the balance of linear momentum

A formal two-scale asymptotic expansion is performed for the displacement u^{ε} , which thus reads

$$\boldsymbol{u}^{\varepsilon}(X,t) = \boldsymbol{u}^{(0)}(X,t) + \sum_{k=1}^{+\infty} \boldsymbol{u}^{(k)}(X,Y,t)\varepsilon^{k}, \qquad (10)$$

where, for all $k \geq 1$, $\boldsymbol{u}^{(k)}$ is periodic with respect to Y. Following [68] we consider the leading order term of the expansion (10) to be independent of the fast variable Y. From formula (4), the expansion (10), and taking into account the property of scale separation, it follows that the deformation gradient tensor can be written as

$$\boldsymbol{F}^{\varepsilon}(X,t) = \sum_{k=0}^{+\infty} \boldsymbol{F}^{(k)}(X,Y,t)\varepsilon^{k}, \qquad (11)$$

349 with the notation

$$\boldsymbol{F}^{(0)} := \boldsymbol{I} + \operatorname{Grad}_{\boldsymbol{X}} \boldsymbol{u}^{(0)} + \operatorname{Grad}_{\boldsymbol{Y}} \boldsymbol{u}^{(1)}, \qquad (12a)$$

$$\boldsymbol{F}^{(k)} := \operatorname{Grad}_{X} \boldsymbol{u}^{(k)} + \operatorname{Grad}_{Y} \boldsymbol{u}^{(k+1)}, \quad \forall \ k \ge 1,$$
(12b)

where Grad_X and Grad_Y are the gradient operators with respect to X and Y, respectively. Now, the following two-scale asymptotic expansion is proposed for the first Piola-Kirchhoff stress tensor T^{ε} ,

$$\boldsymbol{T}^{\varepsilon}(X,t) = \sum_{k=0}^{+\infty} \boldsymbol{T}^{(k)}(X,Y,t)\varepsilon^{k},$$
(13)

where the fields $T^{(k)}$ are periodic with respect to Y. By substituting the power series representation (13) into (8), using the scale separation condition, and multiplying the result by ε , the following multi-scale system is obtained

Div
$$\mathbf{T}^{\varepsilon} = \sum_{k=0}^{+\infty} \mathfrak{D}^{(k)} \varepsilon^k = \mathbf{0},$$
 (14)

356 with

$$\mathfrak{D}^{(0)} := \operatorname{Div}_{Y} \boldsymbol{T}^{(0)}, \tag{15a}$$

$$\mathfrak{D}^{(k)} := \operatorname{Div}_X \boldsymbol{T}^{(k-1)} + \operatorname{Div}_Y \boldsymbol{T}^{(k)}, \quad \forall \ k \ge 1.$$
(15b)

We require that the equilibrium equation (14) is satisfied at every ε , which amounts to impose the conditions

$$\operatorname{Div}_{Y}\boldsymbol{T}^{(0)} = \boldsymbol{0},\tag{16a}$$

$$\operatorname{Div}_X \mathbf{T}^{(k-1)} + \operatorname{Div}_Y \mathbf{T}^{(k)} = \mathbf{0}, \quad \forall \ k \ge 1.$$
 (16b)

³⁵⁹ At this point we introduce the average operator over the microscopic cell, i.e.

$$\langle \bullet \rangle = \frac{1}{|\mathcal{Y}_t|} \int_{\mathcal{Y}_t} \bullet \, \mathrm{d}Y,\tag{17}$$

where $|\mathcal{Y}_t|$ represents the volume of the periodic cell \mathcal{Y}_t at time t. Indeed, because of the deformations and distortions to which the microscopic, reference periodic cell is subjected, \mathcal{Y}_t is different at every time instant. Averaging (16b) over the microscopic cell yields, for k = 1,

$$\langle \operatorname{Div}_X \boldsymbol{T}^{(0)} \rangle + \frac{1}{|\mathcal{Y}_t|} \int_{\partial \mathcal{Y}_t} \boldsymbol{T}^{(1)} \cdot \boldsymbol{N} \mathrm{d}Y = \boldsymbol{0},$$
 (18)

where, on the left-hand side, we have applied the divergence theorem. Since the contributions on the periodic cell boundary $\partial \mathcal{Y}_t$ cancel due to the Yperiodicity, the integral over \mathcal{Y}_t is equal to zero, and (18) becomes

$$\langle \text{Div}_X \boldsymbol{T}^{(0)} \rangle = \boldsymbol{0}.$$
 (19)

Here, we restrict our analysis to the particular case in which the periodic cell can be uniquely chosen independently of X, which implies that the integration over \mathcal{Y}_t and the computation of the divergence commute. This assumption is also referred to as *macroscopic uniformity*, see also [9, 40, 59] for examples dealing with non-macroscopically uniform media in the context of poroelasticity and diffusion. Therefore, Equation (19) can be recast as

$$\operatorname{Div}_X \langle \boldsymbol{T}^{(0)} \rangle = \boldsymbol{0}.$$
 (20)

Equations (16a) and (20) represent, respectively, the local and the homogenized equation associated with the original one, stated in (8). Both equations still need to be supplemented with the corresponding interface, boundary, and initial conditions. Note that, although both problems feature no time derivative, initial conditions are required because $T^{(0)}$ depends on the variable $F_{\rm p}^{(0)}$, which satisfies an evolution equation in time.

We remark that the leading term $\mathbf{T}^{(0)} = \mathbf{T}^{(0)}(X, Y, t)$ of the multi-scale expansion (13) is the unknown, both in (16a) and in (20). To identify $\mathbf{T}^{(0)}$, we propose here to expand $\mathbf{F}_{p}^{\varepsilon}$ and ψ_{ν}^{ε} as

$$\boldsymbol{F}_{\mathrm{p}}^{\varepsilon}(X,t) = \sum_{k=0}^{+\infty} \boldsymbol{F}_{\mathrm{p}}^{(k)}(X,Y,t)\varepsilon^{k}, \qquad (21a)$$

$$\psi_{\nu}^{\varepsilon}(X,t) = \sum_{k=0}^{+\infty} \psi_{\nu}^{(k)}(\boldsymbol{F}_{\mathrm{e}}(X,Y,t),X,Y)\varepsilon^{k}, \qquad (21\mathrm{b})$$

where $\mathbf{F}_{\rm p}^{(k)}$ and $\psi_{\nu}^{(k)}$ are periodic in Y. By using (5), (11) and (21a), we can deduce a series expansion for $\mathbf{F}_{\rm e}^{\varepsilon}$ in powers of ε , where the leading order term $\mathbf{F}_{\rm e}^{(0)}$ is given by

$$\boldsymbol{F}_{e}^{(0)} = \boldsymbol{F}^{(0)} (\boldsymbol{F}_{p}^{(0)})^{-1}.$$
 (22)

Following [15] and [68], $T^{(0)}$ is therefore supplied constitutively as

$$\boldsymbol{T}^{(0)} = J_{\rm p}^{(0)} \frac{\partial \psi_{\nu}^{(0)}}{\partial \boldsymbol{F}_{e}^{(0)}} (\boldsymbol{F}_{\rm p}^{(0)})^{-T}, \qquad (23)$$

with $\psi_{\nu}^{(0)} = \psi_{\nu}^{(0)}(\boldsymbol{F}_{e}^{(0)}(X,Y,t),X,Y)$ and $J_{p}^{(0)} = \det \boldsymbol{F}_{p}^{(0)}$. To obtain the *cell problem*, equation (14) must be supplemented with the corresponding 386 387 interface conditions. This is done by substituting the asymptotic expansions 388 of $\boldsymbol{u}^{\varepsilon}$ and of $\boldsymbol{T}^{\varepsilon}$ into the interface conditions $\llbracket \boldsymbol{u}^{\varepsilon} \rrbracket = \boldsymbol{0}$ and $\llbracket \boldsymbol{T}^{\varepsilon} \cdot \boldsymbol{N}_{\mathcal{Y}} \rrbracket = \boldsymbol{0}$. 389 Both conditions are satisfied at any order of ε . At the order ε^0 , we simply 390 obtain $\llbracket T^{(0)} \cdot N_{\mathcal{V}} \rrbracket = \mathbf{0}$ for the stresses, and that the condition $\llbracket u^{(0)} \rrbracket = \mathbf{0}$ is 391 trivially satisfied, because $\boldsymbol{u}^{(0)}$ depends solely on X and t. Thus, the interface 392 condition on the displacements is written only for $u^{(1)}$ and reads, $[\![u^{(1)}]\!] = 0$. 393 By summarizing these results, the cell problem at zero order of the epsilon 394 parameter can be stated as 395

$$\begin{cases} \operatorname{Div}_{Y} \boldsymbol{T}^{(0)} = \boldsymbol{0}, & \text{in } \mathcal{Y}_{0} \setminus \Gamma_{0} \times \mathcal{T}, \\ [\![\boldsymbol{u}^{(1)}]\!] = \boldsymbol{0}, & \text{on } \Gamma_{0} \times \mathcal{T}, \\ [\![\boldsymbol{T}^{(0)} \cdot \boldsymbol{N}_{\mathcal{Y}}]\!] = \boldsymbol{0}, & \text{on } \Gamma_{0} \times \mathcal{T}. \end{cases}$$
(24)

Together with the cell problem, we also need to formulate the macro-scopic homogenized problem. To this end, we take equation (20) and complete it with a set of boundary conditions. This is done by substituting the asymptotic expansions of T^{ε} and u^{ε} into the boundary conditions $T^{\varepsilon} \cdot N = \bar{T}$ and $u^{\varepsilon} = \bar{u}$, respectively. Thus, equating the coefficients at order ε^0 , and averaging the results over the unit cell, we find the homogenized problem,

$$\begin{cases} \operatorname{Div}_X \langle \boldsymbol{T}^{(0)} \rangle = \boldsymbol{0}, & \operatorname{in} \mathcal{B}_h \times \mathcal{T}, \\ \langle \boldsymbol{T}^{(0)} \rangle \cdot \boldsymbol{N} = \bar{\boldsymbol{T}}, & \operatorname{on} \partial_T \mathcal{B}_h \times \mathcal{T}, \\ \boldsymbol{u}^{(0)} = \bar{\boldsymbol{u}}, & \operatorname{on} \partial_u \mathcal{B}_h \times \mathcal{T}, \end{cases}$$
(25)

where \mathcal{B}_h denotes the homogeneous macro-scale domain in which the homogenized equations are defined.

The problem (25) has to be solved along with a homogenized evolution equation for $\mathbf{F}_{p}^{(0)}$ and the initial condition associated with it. In addition, we remark that, according to (25), the boundary tractions acting on $\partial_T \mathcal{B}_h$ are balanced *only* by the normal component of the average of the leading order stress, $\mathbf{T}^{(0)}$, and *only* the leading order displacement, $\mathbf{u}^{(0)}$, has to be equal to the displacement $\bar{\mathbf{u}}$, imposed on $\partial_u \mathcal{B}_h$.

Remark 2. In the medical scientific literature, there exist studies that iden-410 tify the existence of anatomical boundary layers interposed between the brain 411 surface and tumors (see e.g. [72]). Here we do not address boundary layer 412 phenomena, which are usually neglected in the asymptotic homogenization 413 literature. The homogenization process described in this work is fine for re-414 gions far enough away from the boundary so that its effect is not felt because, 415 close to the boundaries, the material will not behave as an effective material 416 with homogenized coefficients. To properly account for boundary effects, the 417 so-called boundary-layer technique could be used [8, 57]. 418

419 5. Constitutive framework and evolution law

In this section, we prescribe a constitutive equation for the response of the material and, independently, an evolution equation for the tensor of plasticlike distortions.

423 5.1. Constitutive law

In the following, we formulate the local and homogenized problems for a specific constitutive law. In general, this process can be rather cumbersome for complicated strain energy densities, and it becomes even more involved when plastic-like distortions are accounted for. To reduce complexity, we choose a very simple constitutive law for ψ_{ν}^{ε} , such as the De Saint-Venant strain energy density,

$$\psi_{\nu}^{\varepsilon} = \frac{1}{2} \boldsymbol{E}_{e}^{\varepsilon} : \mathscr{C}^{\varepsilon} : \boldsymbol{E}_{e}^{\varepsilon}, \qquad (26)$$

where $\boldsymbol{E}_{e}^{\varepsilon} = \frac{1}{2} \left((\boldsymbol{F}_{e}^{\varepsilon})^{T} \boldsymbol{F}_{e}^{\varepsilon} - \boldsymbol{I} \right)$ is the elastic Green-Lagrange strain tensor and $\mathscr{C}^{\varepsilon}(X) = \mathscr{C}(X, Y)$ is the positive definite fourth-order elasticity tensor, which satisfies both major and minor symmetries, i.e. $\mathscr{C}_{ijkl} = \mathscr{C}_{jikl} = \mathscr{C}_{ijlk} = \mathscr{C}_{klij}$. Particularly, we consider that the constituents of the heterogeneous material are isotropic, and thus

$$\mathscr{C}^{\varepsilon} = 3\kappa^{\varepsilon}\mathscr{K} + 2\mu^{\varepsilon}\mathscr{M}, \qquad (27)$$

where $\kappa^{\varepsilon}(X) = \kappa(X, Y)$ is the bulk modulus, $\mu^{\varepsilon}(X) = \mu(X, Y)$ is the shear 435 modulus, and the fourth-order tensors $\mathscr{K} = \frac{1}{3}(\mathbf{I} \otimes \mathbf{I})$ and $\mathscr{M} = \mathscr{I} - \mathscr{K}$ 436 extract the spherical and the deviatoric part, respectively, of a symmetric 437 second-order tensor A, i.e., $\mathscr{K} : A = \frac{1}{3} \operatorname{tr}(A) I$ and $\mathscr{M} : A = A - \frac{1}{3} \operatorname{tr}(A) I :=$ 438 $\operatorname{dev}(\mathbf{A})$ [84, 85]. We remark that the fourth-order identity tensor \mathscr{I} is the 439 identity operator over the linear subspace of symmetric second-order tensors. 440 Indeed, for every A such that $A = A^T$, it holds that $\mathscr{I} : A = A$. In 441 terms of I, an explicit expression of \mathscr{I} is given by $\mathscr{I} = \frac{1}{2} [I \otimes I + I \otimes I]$ (in 442 components: $\mathscr{I}_{ijkl} = \frac{1}{2} [I_{ik}I_{jl} + I_{il}I_{jk}] [17]).$ 443

We can identify the leading order term in the expansion of the constitutive law (26), which reads

$$\psi_{\nu}^{(0)} = \frac{1}{2} \boldsymbol{E}_{e}^{(0)} : \mathscr{C} : \boldsymbol{E}_{e}^{(0)}, \qquad (28)$$

with $\boldsymbol{E}_{e}^{(0)} = \frac{1}{2} \left((\boldsymbol{F}_{e}^{(0)})^{T} \boldsymbol{F}_{e}^{(0)} - \boldsymbol{I} \right)$. We recall that, although the expression of $\psi_{\nu}^{(0)}$ in (28) depends only on $\boldsymbol{E}_{e}^{(0)}$, the material coefficient \mathscr{C} is still a twoscale function and should be thus interpreted as $\mathscr{C}(X,Y)$. As a consequence, $\psi_{\nu}^{(0)}$ is not homogenized yet.

 $_{450}$ By taking into account the major and minor symmetries of \mathscr{C} , we obtain

$$\boldsymbol{S}_{\nu}^{(0)} = \frac{\partial \psi_{\nu}^{(0)}}{\partial \boldsymbol{E}_{e}^{(0)}} = \mathscr{C} : \boldsymbol{E}_{e}^{(0)} = \lambda \operatorname{tr}(\boldsymbol{E}_{e}^{(0)})\boldsymbol{I} + 2\mu \boldsymbol{E}_{e}^{(0)}, \qquad (29)$$

where $\mathbf{S}_{\nu}^{(0)}$ is the leading order term of the second Piola-Kirchhoff stress tensor written with respect to the natural state, $\lambda = \kappa - \frac{2}{3}\mu$ is Lamé's constant, and $\mathbf{E}_{e}^{(0)}$ is given by

$$\boldsymbol{E}_{e}^{(0)} = (\boldsymbol{F}_{p}^{(0)})^{-T} \left(\boldsymbol{E}^{(0)} - \boldsymbol{E}_{p}^{(0)} \right) (\boldsymbol{F}_{p}^{(0)})^{-1},$$
(30)

with $\boldsymbol{E}^{(0)} = \frac{1}{2} \left((\boldsymbol{F}^{(0)})^T \boldsymbol{F}^{(0)} - \boldsymbol{I} \right)$ and $\boldsymbol{E}_{p}^{(0)} = \frac{1}{2} \left((\boldsymbol{F}_{p}^{(0)})^T \boldsymbol{F}_{p}^{(0)} - \boldsymbol{I} \right)$.

⁴⁵⁵ By pulling $S_{\nu}^{(0)}$ back to the reference configuration, and recalling that the ⁴⁵⁶ plastic-like distortions are assumed to be isochoric in our framework, (i.e. ⁴⁵⁷ $J_{\rm p}^{\varepsilon} = 1$), we obtain the second Piola-Kirchhoff stress tensor

$$\boldsymbol{S}^{(0)} = \mathscr{C}_{\mathrm{R}} : (\boldsymbol{E}^{(0)} - \boldsymbol{E}_{\mathrm{p}}^{(0)}), \qquad (31)$$

458 where

$$\mathscr{C}_{\mathrm{R}} = (\boldsymbol{F}_{\mathrm{p}}^{(0)})^{-1} \underline{\otimes} (\boldsymbol{F}_{\mathrm{p}}^{(0)})^{-1} : \mathscr{C} : (\boldsymbol{F}_{\mathrm{p}}^{(0)})^{-T} \underline{\otimes} (\boldsymbol{F}_{\mathrm{p}}^{(0)})^{-T}$$
$$= 3\lambda \mathscr{K}_{\mathrm{p}}^{(0)} + 2\mu \mathscr{I}_{\mathrm{p}}^{(0)}, \qquad (32)$$

is the elasticity tensor pulled-back to the reference configuration through $F_{\rm p}^{(0)}$, and, upon setting $B_{\rm p}^{(0)} = (F_{\rm p}^{(0)})^{-1} (F_{\rm p}^{(0)})^{-T}$, we employed the notation

$$\mathscr{K}_{\rm p}^{(0)} = \frac{1}{3} \mathscr{B}_{\rm p}^{(0)} \otimes \mathscr{B}_{\rm p}^{(0)},$$
 (33a)

$$\mathscr{I}_{\mathrm{p}}^{(0)} = \frac{1}{2} \left[\boldsymbol{B}_{\mathrm{p}}^{(0)} \underline{\otimes} \boldsymbol{B}_{\mathrm{p}}^{(0)} + \boldsymbol{B}_{\mathrm{p}}^{(0)} \overline{\otimes} \boldsymbol{B}_{\mathrm{p}}^{(0)} \right].$$
(33b)

We remark that $\mathscr{K}_{p}^{(0)}$ extracts the "volumetric part" of a generic secondorder tensor, taken with respect to the inverse plastic metric tensor $\boldsymbol{B}_{p}^{(0)}$ i.e. for all $\boldsymbol{A} = \boldsymbol{A}^{T}$, it holds that $\mathscr{K}_{p}^{(0)} : \boldsymbol{A} = \frac{1}{3} tr(\boldsymbol{B}_{p}^{(0)} \boldsymbol{A}) \boldsymbol{B}_{p}^{(0)}$. Furthermore, $\mathscr{I}_{p}^{(0)}$ transforms \boldsymbol{A} into $\mathscr{I}_{p}^{(0)} : \boldsymbol{A} = \boldsymbol{B}_{p}^{(0)} \boldsymbol{A} \boldsymbol{B}_{p}^{(0)}$ and $\mathscr{M}_{p}^{(0)} = \mathscr{I}_{p}^{(0)} - \mathscr{K}_{p}^{(0)}$ extracts the "deviatoric part" of \boldsymbol{A} with respect to the metric tensor $\boldsymbol{B}_{p}^{(0)}$, i.e. $\mathscr{M}_{p}^{(0)} : \boldsymbol{A} = \boldsymbol{B}_{p}^{(0)} \boldsymbol{A} \boldsymbol{B}_{p}^{(0)} - \frac{1}{3} tr(\boldsymbol{B}_{p}^{(0)} \boldsymbol{A}) \boldsymbol{B}_{p}^{(0)}$. We note that similar results have been obtained in the case of non-linear elasticity in [25]. Next, we notice that $\boldsymbol{F}^{(0)}$ can be written as

$$\boldsymbol{F}^{(0)} = \boldsymbol{I} + \boldsymbol{H},\tag{34}$$

with $\boldsymbol{H} = \operatorname{Grad}_{X}\boldsymbol{u}^{(0)} + \operatorname{Grad}_{Y}\boldsymbol{u}^{(1)}$. Thus, by substituting (34) in $\boldsymbol{E}_{e}^{(0)}$, the result into (31), and retaining only the terms linear in \boldsymbol{H} , $\boldsymbol{S}^{(0)}$ can be linearized as

$$\boldsymbol{S}_{\text{lin}}^{(0)} = \mathscr{C}_{\text{R}} : (\text{sym}\boldsymbol{H} - \boldsymbol{E}_{\text{p}}^{(0)}). \tag{35}$$

We recall now that, at the leading order, the first Piola-Kirchhoff stress tensor reads $T^{(0)} = F^{(0)}S^{(0)}$. Hence, its linearized form is given by

$$\boldsymbol{T}_{\text{lin}}^{(0)} = \mathscr{C}_{\text{R}} : \text{sym}\boldsymbol{H} - (\boldsymbol{I} + \boldsymbol{H})(\mathscr{C}_{\text{R}} : \boldsymbol{E}_{\text{p}}^{(0)}).$$
(36)

Looking at the definition of \mathscr{C}_{R} in (32), it can be noticed that our model re-474 solves at the macro-scale the structural evolution of the considered medium 475 through the dependence of \mathscr{C}_{R} on $\boldsymbol{F}_{\mathrm{p}}^{(0)}$, which indeed describes the produc-476 tion of material inhomogeneities [21, 22, 23]. Additionally, our model is also 477 capable of simultaneously resolving the material heterogeneities at both the 478 micro- and macro-scale through the dependence of $\mathscr{C}_{\mathbf{R}}$ on X and Y. The lat-479 ter dependence in fact, keeps track of the variability of the elastic coefficient 480 at both scales. 481

Because of Equations (33a) and (33b), \mathscr{C}_{R} possesses the same symmetry properties of \mathscr{C} , i.e.

$$(\mathscr{C}_{\mathbf{R}})_{IJKL} = (\mathscr{C}_{\mathbf{R}})_{JIKL} = (\mathscr{C}_{\mathbf{R}})_{IJLK} = (\mathscr{C}_{\mathbf{R}})_{KLIJ},$$
(37)

484 and therefore, $oldsymbol{T}_{ ext{lin}}^{(0)}$ can be written as

$$\boldsymbol{T}_{\text{lin}}^{(0)} = \mathscr{C}_{\text{R}} : \boldsymbol{H} - (\boldsymbol{I} + \boldsymbol{H})(\mathscr{C}_{\text{R}} : \boldsymbol{E}_{\text{p}}^{(0)}).$$
(38)

Local problem. Substituting (38) in the equation of the local problem (24), the linear momentum balance law is rephrased as

$$\operatorname{Div}_{Y}\left[\mathscr{C}_{\mathrm{R}}:\boldsymbol{H}-(\boldsymbol{I}+\boldsymbol{H})(\mathscr{C}_{\mathrm{R}}:\boldsymbol{E}_{\mathrm{p}}^{(0)})\right]=\boldsymbol{0},\tag{39}$$

487 or, equivalently,

$$\operatorname{Div}_{Y} \left[\mathscr{C}_{\mathrm{R}} : \operatorname{Grad}_{Y} \boldsymbol{u}^{(1)} - \operatorname{Grad}_{Y} \boldsymbol{u}^{(1)} (\mathscr{C}_{\mathrm{R}} : \boldsymbol{E}_{\mathrm{p}}^{(0)}) \right] = -\operatorname{Div}_{Y} \left[\mathscr{C}_{\mathrm{R}} : \operatorname{Grad}_{X} \boldsymbol{u}^{(0)} - (\boldsymbol{I} + \operatorname{Grad}_{X} \boldsymbol{u}^{(0)}) (\mathscr{C}_{\mathrm{R}} : \boldsymbol{E}_{\mathrm{p}}^{(0)}) \right].$$
(40)

In the absence of plastic distortions, i.e., when $F_{\rm p}^{\varepsilon} = I$, Equation (40) coincides with the equation of the classical cell problem encountered in the homogeneization of linear elasticity, which is known to admit a unique solution, up to a Y-constant function, if the average over the cell of the right-hand-side vanishes identically (in the jargon of Homogenization Theory, this condition is referred to as solvability condition or compatibility condition) [5]. In our ⁴⁹⁴ case, since the pulled-back elasticity tensor $\mathscr{C}_{\mathbf{R}}$ is periodic in Y, while $\boldsymbol{u}^{(0)}$ is ⁴⁹⁵ independent of Y, the solvability condition is satisfied, i.e.,

$$\left\langle \operatorname{Div}_{Y}\left[\mathscr{C}_{\mathrm{R}}:\operatorname{Grad}_{X}\boldsymbol{u}^{(0)}-(\boldsymbol{I}+\operatorname{Grad}_{X}\boldsymbol{u}^{(0)})(\mathscr{C}_{\mathrm{R}}:\boldsymbol{E}_{\mathrm{p}}^{(0)})\right]\right\rangle=\boldsymbol{0}.$$
 (41)

Exploiting the linearity of equation (40) in $\boldsymbol{u}^{(1)}$, we make the ansatz

$$\boldsymbol{u}^{(1)}(X,Y,t) = \boldsymbol{\xi}(X,Y,t) : \operatorname{Grad}_{X} \boldsymbol{u}^{(0)}(X,t) + \boldsymbol{\omega}(X,Y,t),$$
(42)

where $\boldsymbol{\xi}$ and $\boldsymbol{\omega}$ are a third-order tensor field and a vector field, both periodic in Y.

We now require that $\boldsymbol{\xi}$ and $\boldsymbol{\omega}$ satisfy two independent cell problems. The cell problem for $\boldsymbol{\xi}$ reads

$$\begin{cases} \operatorname{Div}_{Y} \left[\mathscr{C}_{\mathrm{R}} : T\operatorname{Grad}_{Y} \boldsymbol{\xi} - T\operatorname{Grad}_{Y} \boldsymbol{\xi} (\mathscr{C}_{\mathrm{R}} : \boldsymbol{E}_{\mathrm{p}}^{(0)}) \right] \\ = \operatorname{Div}_{Y} \left[- \mathscr{C}_{\mathrm{R}} + \boldsymbol{I} \underline{\otimes} (\mathscr{C}_{\mathrm{R}} : \boldsymbol{E}_{\mathrm{p}}^{(0)}) \right], & \text{in } \mathcal{Y}_{0} \setminus \Gamma_{0} \times \mathcal{T}, \\ \left[\boldsymbol{\xi} \right] = \boldsymbol{0}, & \text{on } \Gamma_{0} \times \mathcal{T}, \\ \left[\left[\mathscr{C}_{\mathrm{R}} : T\operatorname{Grad}_{Y} \boldsymbol{\xi} - T\operatorname{Grad}_{Y} \boldsymbol{\xi} (\mathscr{C}_{\mathrm{R}} : \boldsymbol{E}_{\mathrm{p}}^{(0)}) \right] \\ + \mathscr{C}_{\mathrm{R}} - \boldsymbol{I} \underline{\otimes} (\mathscr{C}_{\mathrm{R}} : \boldsymbol{E}_{\mathrm{p}}^{(0)}) \right] \cdot \boldsymbol{N}_{\mathcal{Y}} \right] = \boldsymbol{0}, & \text{on } \Gamma_{0} \times \mathcal{T}. \end{cases}$$

$$(43)$$

⁵⁰¹ Before going further, some words of explanation on the notation are nec-⁵⁰² essary. First, we notice that $\operatorname{Grad}_Y \boldsymbol{\xi}$ is a fourth-order tensor function, which ⁵⁰³ admits the representation $\operatorname{Grad}_Y \boldsymbol{\xi} = (\partial \xi_{ABC})/(\partial Y_D) \boldsymbol{e}_A \otimes \boldsymbol{e}_B \otimes \boldsymbol{e}_C \otimes \boldsymbol{e}_D$. Then, ⁵⁰⁴ $T\operatorname{Grad}_Y \boldsymbol{\xi}$ is a fourth-order tensor function obtained by ordering the indices ⁵⁰⁵ of $\operatorname{Grad}_Y \boldsymbol{\xi}$ in the following fashion

$$T \operatorname{Grad}_{Y} \boldsymbol{\xi} = (T \operatorname{Grad}_{Y} \boldsymbol{\xi})_{ABCD} \boldsymbol{e}_{A} \otimes \boldsymbol{e}_{B} \otimes \boldsymbol{e}_{C} \otimes \boldsymbol{e}_{D}$$
$$= (\operatorname{Grad}_{Y} \boldsymbol{\xi})_{ACDB} \boldsymbol{e}_{A} \otimes \boldsymbol{e}_{B} \otimes \boldsymbol{e}_{C} \otimes \boldsymbol{e}_{D}$$
$$= \frac{\partial \xi_{ACD}}{\partial Y_{B}} \boldsymbol{e}_{A} \otimes \boldsymbol{e}_{B} \otimes \boldsymbol{e}_{C} \otimes \boldsymbol{e}_{D}.$$
(44)

The cell problem for $\boldsymbol{\omega}$ is given by

$$\begin{cases} \operatorname{Div}_{Y} \left[\mathscr{C}_{\mathrm{R}} : \operatorname{Grad}_{Y} \boldsymbol{\omega} - \operatorname{Grad}_{Y} \boldsymbol{\omega} (\mathscr{C}_{\mathrm{R}} : \boldsymbol{E}_{\mathrm{p}}^{(0)}) \right] \\ = \operatorname{Div}_{Y} \left[\mathscr{C}_{\mathrm{R}} : \boldsymbol{E}_{\mathrm{p}}^{(0)} \right], & \text{in } \mathcal{Y}_{0} \setminus \Gamma_{0} \times \mathcal{T}, \\ \left[\boldsymbol{\omega} \right] = \boldsymbol{0}, & \text{on } \Gamma_{0} \times \mathcal{T}, \\ \left[\left(\mathscr{C}_{\mathrm{R}} : \operatorname{Grad}_{Y} \boldsymbol{\omega} - \operatorname{Grad}_{Y} \boldsymbol{\omega} (\mathscr{C}_{\mathrm{R}} : \boldsymbol{E}_{\mathrm{p}}^{(0)}) \right. \\ \left. - \mathscr{C}_{\mathrm{R}} : \boldsymbol{E}_{\mathrm{p}}^{(0)} \right) \cdot \boldsymbol{N}_{\mathcal{Y}} \right] = \boldsymbol{0}, & \text{on } \Gamma_{0} \times \mathcal{T}. \end{cases}$$
(45)

By virtue of the linearization process, we obtain two auxiliary cell problems 507 where the macroscopic term $\operatorname{Grad}_X \boldsymbol{u}^{(0)}$ is not explicitly present. Indeed, this 508 is in general possible only when accounting for the linearized deformations' 509 regime, see also [15]. Then, the dependence of the macro-scale variable is 510 given through the tensor $F_{\rm p}^{(0)}$, which describes the plastic-like distortions. 511 Moreover, if $\boldsymbol{F}_{p}^{(0)}$ only depends on time, as is the case in [2], the cell problems 512 are also decoupled in the spatial micro- and macro-variables provided that the 513 elasticity tensor solely depends on the microscale variable. The cell problems 514 are in any case time-dependent, as they encode the evolution of the material 515 response and its link with the plastic-like distortions. 516

⁵¹⁷ Homogenized problem. From (36) and (42), the homogenized problem rewrites

$$\begin{cases} \operatorname{Div}_{X} \left[\hat{\mathscr{C}}_{\mathrm{R}} : \operatorname{Grad}_{X} \boldsymbol{u}^{(0)} \right] = -\operatorname{Div}_{X} \left[\hat{\boldsymbol{D}}_{\mathrm{R}} \right], & \text{in } \mathcal{B}_{h} \times \mathcal{T}, \\ (\hat{\mathscr{C}}_{\mathrm{R}} : \operatorname{Grad}_{X} \boldsymbol{u}^{(0)}) \cdot \boldsymbol{N} + \hat{\boldsymbol{D}}_{\mathrm{R}} \cdot \boldsymbol{N} = \bar{\boldsymbol{T}}, & \text{on } \partial_{T} \mathcal{B}_{h} \times \mathcal{T}, \\ \boldsymbol{u}^{(0)} = \bar{\boldsymbol{u}}, & \text{on } \partial_{u} \mathcal{B}_{h} \times \mathcal{T}, \end{cases}$$
(46)

518 where

$$\hat{\mathscr{C}}_{\mathrm{R}} = \left\langle \mathscr{C}_{\mathrm{R}} + \mathscr{C}_{\mathrm{R}} : T \operatorname{Grad}_{Y} \boldsymbol{\xi} - T \operatorname{Grad}_{Y} \boldsymbol{\xi} (\mathscr{C}_{\mathrm{R}} : \boldsymbol{E}_{\mathrm{p}}^{(0)}) - \boldsymbol{I} \underline{\otimes} (\mathscr{C}_{\mathrm{R}} : \boldsymbol{E}_{\mathrm{p}}^{(0)}) \right\rangle, \quad (47a)$$
$$\hat{\boldsymbol{D}}_{\mathrm{R}} = \left\langle \mathscr{C}_{\mathrm{R}} : \operatorname{Grad}_{Y} \boldsymbol{\omega} - \operatorname{Grad}_{Y} \boldsymbol{\omega} (\mathscr{C}_{\mathrm{R}} : \boldsymbol{E}_{\mathrm{p}}^{(0)}) - \mathscr{C}_{\mathrm{R}} : \boldsymbol{E}_{\mathrm{p}}^{(0)} \right\rangle. \quad (47b)$$

Remark 3. In the absence of distortions, that is for $F_{\rm p}^{\varepsilon} = I$, the cell problems (43) and (45) reduce to one single cell problem,

$$\begin{cases} \operatorname{Div}_{Y}[\mathscr{C} + \mathscr{C} : T\operatorname{Grad}_{Y}\boldsymbol{\xi}] = \boldsymbol{0}, & \text{in } \mathcal{Y}_{0} \setminus \Gamma_{0} \times \mathcal{T}, \\ \llbracket \boldsymbol{\xi} \rrbracket = \boldsymbol{0}, & \text{on } \Gamma_{0} \times \mathcal{T}, \\ \llbracket (\mathscr{C} + \mathscr{C} : T\operatorname{Grad}_{Y}\boldsymbol{\xi}) \cdot \boldsymbol{N}_{\mathcal{Y}} \rrbracket = \boldsymbol{0}, & \text{on } \Gamma_{0} \times \mathcal{T}. \end{cases}$$
(48)

This is due to the fact that the symmetric tensor $\boldsymbol{E}_{p}^{(0)}$ appearing in (40) is equal to zero. On the other hand, the homogenized problem is rewritten as follows,

$$\begin{cases} \operatorname{Div}_{X}[\hat{\mathscr{C}}:\operatorname{Grad}_{X}\boldsymbol{u}^{(0)}] = \boldsymbol{0}, & in \ \mathcal{B}_{h} \times \mathcal{T}, \\ (\hat{\mathscr{C}}:\operatorname{Grad}_{X}\boldsymbol{u}^{(0)}) \cdot \boldsymbol{N} = \bar{\boldsymbol{T}}, & on \ \partial_{T}\mathcal{B}_{h} \times \mathcal{T}, \\ \boldsymbol{u}^{(0)} = \bar{\boldsymbol{u}}, & on \ \partial_{u}\mathcal{B}_{h} \times \mathcal{T}, \end{cases}$$
(49)

where $\hat{\mathscr{C}} = \langle \mathscr{C} + \mathscr{C} : T \operatorname{Grad}_Y \boldsymbol{\xi} \rangle$ is the effective elasticity tensor. Formulations (48) and (49) are the counterparts of (24) and (25), respectively, when plastic-like distortions are neglected and a linearized approach for the deformations is considered. Particularly, (48) and (49) identify identically with classical results in the asymptotic homogenization literature [5, 77].

529 5.2. Evolution law

Several procedures can be adopted to establish a proper evolution law 530 for the inelastic distortions. One choice is to follow a phenomenological 531 approach, which should be based on experimental evidences and comply with 532 suitable constitutive requirements [29]. On the other hand, one could invoke 533 some general principles, such as the invariance of the evolution law with 534 respect to a class of transformations and thermodynamic constraints [21, 22, 535 23. Within the latter approach, and adapting the theoretical framework 536 explored in [21, 22, 23, 29], an evolution equation for the inelastic distortions 537 has been studied in [19]. Therein, the plastic-like distortions describe a 538 remodeling process with the following assumptions: (i) $F_{\rm p}$ is restricted by the 539 constraint $J_{\rm p} = 1$, (ii) the solid phase exhibits hyperelastic behavior, and (iii) 540 the considered system remodels when the stress induced by external loading 541 exceeds a characteristic threshold. An evolution law for $F_{\rm p}$ satisfying these 542 conditions, and compatible with the Dissipation inequality [12, 32, 33, 34], 543 is given by 544

sym
$$\left(\boldsymbol{C} \boldsymbol{F}_{\mathrm{p}}^{-1} \dot{\boldsymbol{F}}_{\mathrm{p}} \right) = \gamma \left[\| \operatorname{dev} \boldsymbol{\sigma} \| - \sqrt{\frac{2}{3}} \sigma_{y} \right]_{+} \frac{\operatorname{dev}(\boldsymbol{\Sigma}) \boldsymbol{C}}{\| \operatorname{dev} \boldsymbol{\sigma} \|},$$
 (50)

where $\boldsymbol{\sigma}$ is the Cauchy stress tensor, $\operatorname{dev}(\boldsymbol{\Sigma}) = \boldsymbol{\Sigma} - \frac{1}{3} \operatorname{tr}(\boldsymbol{\Sigma}) \boldsymbol{I}$, is the deviatoric 545 part of the Mandell stress tensor $\Sigma = CS$ being the Mandel stress tensor, 546 and $\boldsymbol{S} = \boldsymbol{F}^{-1}\boldsymbol{T}$ the second Piola-Kirchhoff stress tensor. Moreover, γ is a 547 strictly positive model parameter, $\sigma_y > 0$ is the yield, or threshold, stress, 548 and the operator $[A]_+$ is such that, for any real number A, $[A]_+ = A$, if A > 0, 549 and $[A]_{+} = 0$ otherwise. As anticipated in the Introduction, in the present 550 context the physical meaning of the plastic-like distortions, represented by 551 $F_{\rm p}$, is that of structural reorganization, i.e. remodeling, as is the case in 552 biological tissues when the adhesion bonds among cells or the structure of 553 the ECM reorganize themselves. 554

Although Equation (50) has been successfully used to describe some biological situations in which the onset of remodeling is subordinated to the excess of the yield stress σ_y , the homogenization of the evolution law (50) is too complicated. For this reason, in this work, we replace (50) with a much easier law of the type

sym
$$\left(\boldsymbol{C}(\boldsymbol{F}_{\mathrm{p}})^{-1} \dot{\boldsymbol{F}}_{\mathrm{p}} \right) = \gamma \operatorname{dev}(\boldsymbol{\Sigma}) \boldsymbol{C},$$
 (51)

according to which no stress-activation criterion is supplied. Clearly, this choice may turn out to be unrealistic in many circumstances, but it can still be useful to understand the essence of some stress-driven remodeling processes.

We need to clarify that, although in some sentences of this work we 564 mentioned growth, our model focuses on *pure* remodeling. This is reflected 565 by the condition det $\boldsymbol{F}_{\rm p} = 1$, and, more importantly, by the fact that the 566 evolution laws (50) and (51) are triggered and controlled exclusively by me-567 chanical factors. On the one hand, the requirement $\det F_{\rm p} = 1$ means that 568 the plastic-like distortions are isochoric and, thus, unable to describe volu-569 metric growth. On the other hand, the evolution laws for \boldsymbol{F}_{p} , i.e., Eqs. (50) 570 or (51), imply that remodeling is viewed as a consequence of the mechanical 571 environment only: When mechanical stress exceeds a given threshold (see 572 also [29, 34]), the internal structure of the tissue starts to vary. In other 573 words, in the present framework, no biochemical phenomena are accounted 574 for as possible activators of remodeling. This is a remarkable difference with 575 growth, which, in contrast, occurs only when the concentration of nutrients 576 is above a certain threshold value [2, 10, 3, 26, 52]. Our results do not apply 577 to growth as they stand, nonetheless, the theory can be adapted to model 578 growth by doing some necessary modifications. This is the reason why in 579 the abstract we stated that our study offers "a robust framework that can be 580 readily generalized to growth and remodeling of nonlinear composites". 581

To homogenize (51), the first step is to rewrite it as

sym
$$\left(\boldsymbol{C}^{\varepsilon} (\boldsymbol{F}_{p}^{\varepsilon})^{-1} \dot{\boldsymbol{F}}_{p}^{\varepsilon} \right) = \gamma^{\varepsilon} dev(\boldsymbol{\Sigma}^{\varepsilon}) \boldsymbol{C}^{\varepsilon},$$
 (52)

by admitting that $\gamma^{\varepsilon}(X) = \gamma(X, Y)$ is a rapidly oscillating strictly positive function. Moreover, by performing the power expansion for Σ^{ε} ,

$$\Sigma^{\varepsilon}(X,t) = \sum_{k=0}^{+\infty} \Sigma^{(k)}(X,Y,t)\varepsilon^k, \qquad (53)$$

and using (31), the leading order term of Σ^{ε} is

$$\boldsymbol{\Sigma}^{(0)} = \boldsymbol{C}^{(0)} \big[\mathscr{C}_{\mathrm{R}} : (\boldsymbol{E}^{(0)} - \boldsymbol{E}_{\mathrm{p}}^{(0)}) \big].$$
 (54)

In the limit of small elastic deformations, in (54) we must neglect non-linear terms in H. Therefore, $\Sigma^{(0)}$ is approximated with

$$\boldsymbol{\Sigma}_{\text{lin}}^{(0)} = \mathscr{C}_{\text{R}} : \text{sym}\boldsymbol{H} - (\boldsymbol{I} + 2\text{sym}\boldsymbol{H})(\mathscr{C}_{\text{R}} : \boldsymbol{E}_{\text{p}}^{(0)}).$$

⁵⁸⁸ By virtue of (12a), symH splits additively as the sum of

$$\operatorname{sym} \boldsymbol{H} = \boldsymbol{E}_X^{(0)} + \boldsymbol{E}_Y^{(1)}, \qquad (55)$$

⁵⁸⁹ where, for k = 0, 1, and $j_k = X, Y$,

$$\boldsymbol{E}_{j_k}^{(k)} = \frac{1}{2} \left[\operatorname{Grad}_{j_k} \boldsymbol{u}^{(k)} + (\operatorname{Grad}_{j_k} \boldsymbol{u}^{(k)})^T \right].$$
(56)

⁵⁹⁰ By using (55) and (42), we can now rewrite $\Sigma_{\text{lin}}^{(0)}$ as

$$\boldsymbol{\Sigma}_{\text{lin}}^{(0)} = \mathscr{A}_{\text{R}} : \text{Grad}_{X} \boldsymbol{u}^{(0)} + \mathscr{B}_{\text{R}} : \text{Grad}_{Y} \boldsymbol{\omega} - \mathscr{C}_{\text{R}} : \boldsymbol{E}_{\text{p}}^{(0)},$$
(57)

591 with

$$\mathcal{A}_{\mathrm{R}} = \mathscr{C}_{\mathrm{R}} + \mathscr{C}_{\mathrm{R}} : T\mathrm{Grad}_{Y}\boldsymbol{\xi} - \boldsymbol{I}\overline{\otimes}(\mathscr{C}_{\mathrm{R}} : \boldsymbol{E}_{\mathrm{p}}^{(0)}) + \left[\boldsymbol{I}\underline{\otimes}(\mathscr{C}_{\mathrm{R}} : \boldsymbol{E}_{\mathrm{p}}^{(0)})\right] : \left[T\mathrm{Grad}_{Y}\boldsymbol{\xi} + {}^{t}(T\mathrm{Grad}_{Y}\boldsymbol{\xi})\right],$$
(58a)

$$\mathscr{B}_{\mathrm{R}} = \mathscr{C}_{\mathrm{R}} + \boldsymbol{I} \overline{\underline{\otimes}} (\mathscr{C}_{\mathrm{R}} : \boldsymbol{E}_{\mathrm{p}}^{(0)}).$$
 (58b)

In Equation (58a), the symbol ${}^{t}(\bullet)$ transposes the fourth-order tensor to which it is applied by exchanging the order of its first pair of indices only, i.e., given an arbitrary fourth-order tensor $\mathscr{T} = \mathscr{T}_{ABCD} \boldsymbol{e}_{A} \otimes \boldsymbol{e}_{B} \otimes \boldsymbol{e}_{C} \otimes \boldsymbol{e}_{D}$, ${}^{t}\mathscr{T}$ reads

$${}^{t}\mathscr{T} = \mathscr{T}_{BACD} \boldsymbol{e}_{A} \otimes \boldsymbol{e}_{B} \otimes \boldsymbol{e}_{C} \otimes \boldsymbol{e}_{D}.$$

$$(59)$$

Note that in the calculations performed to obtain \mathscr{A}_{R} and \mathscr{B}_{R} in (57), we employed the following properties: given two second-order tensors \boldsymbol{A} and \boldsymbol{U} , with \boldsymbol{A} being symmetric, it holds that

$$\boldsymbol{U}\boldsymbol{A} = (\boldsymbol{I}\underline{\otimes}\boldsymbol{A}): \boldsymbol{U},\tag{60a}$$

$$\boldsymbol{U}^T \boldsymbol{A} = (\boldsymbol{I} \overline{\otimes} \boldsymbol{A}) : \boldsymbol{U}. \tag{60b}$$

⁵⁹⁹ Finally, by substituting the expansions of Σ^{ε} and F_{p}^{ε} in (52), equating ⁶⁰⁰ the leading order terms, excluding non-linear terms of H and averaging, the ⁶⁰¹ homogenized evolution law for the plastic-like distortions is

$$\operatorname{sym}\left[\langle \boldsymbol{C}_{\operatorname{lin}}^{(0)}(\boldsymbol{F}_{\operatorname{p}}^{(0)})^{-1}\overline{\boldsymbol{F}_{\operatorname{p}}^{(0)}}\rangle\right] = -\langle \gamma \operatorname{dev}(\boldsymbol{\Sigma}_{\operatorname{lin}}^{(0)})\rangle - \langle \gamma(\mathscr{C}_{\operatorname{R}}:\boldsymbol{E}_{\operatorname{p}}^{(0)})(\boldsymbol{C}_{\operatorname{lin}}^{(0)}-\boldsymbol{I})\rangle, (61)$$

⁶⁰² where $\Sigma_{\text{lin}}^{(0)}$ is given in (57) and

$$\boldsymbol{C}_{\text{lin}}^{(0)} = \boldsymbol{I} + 2\text{sym}\boldsymbol{H}$$

= $\boldsymbol{I} + 2(\mathscr{I} + \mathscr{I} : T\text{Grad}_{Y}\boldsymbol{\xi}) : \text{Grad}_{X}\boldsymbol{u}^{(0)} + 2\mathscr{I} : \text{Grad}_{Y}\boldsymbol{\omega}.$ (62)

⁶⁰³ We note that, to compute $C_{\text{lin}}^{(0)}$, we must first determine $\boldsymbol{\xi}$ and $\boldsymbol{\omega}$, which is ⁶⁰⁴ done by solving the local problems (43) and (45). Furthermore, Equation ⁶⁰⁵ (61) needs to be supplemented with an initial condition for $\boldsymbol{F}_{p}^{(0)}$.

Remark 4. In the linearized theory of elasticity, even when the individual 606 constituents of a given composite material are isotropic, the effective elas-607 tic coefficients may turn out to be anisotropic, depending on the geometric 608 properties of the micro-structure. In fact, when the Homogenization Theory 609 is applied, the anisotropy arises quite naturally due to the solution of the 610 local cell problems [5, 8]. In fact, the homogenized material is anisotropic 611 also in the case of rather simple cells, see for instance [61], where an ex-612 plicit deviation-from- isotropy function is introduced in the context of cubic 613 symmetric elasticity tensors arising from asympototic homogenization. This 614 has noticeable repercussions also on the evolution law that should be chosen 615 for a correct description of remodeling. To see this, we first notice that, for 616 an isotropic medium, the evolution law of the plastic-like distortions can be 617 formulated in terms of tensor $B_{\rm p}$, since the constitutive framework is such 618 that \boldsymbol{F}_{p} does not feature explicitly in any constitutive function (see e.g. [78]). 619 In such cases, a possible evolution law for B_{p} may be given in the form 620

$$\dot{\boldsymbol{B}}_{\mathrm{p}} = \gamma \boldsymbol{B}_{\mathrm{p}} \mathrm{dev}(\boldsymbol{\Sigma}). \tag{63}$$

Equation (63) is, in fact, in harmony with the symmetry properties of the material Mandel stress tensor, Σ , i.e., $B_{\rm p}\Sigma = (B_{\rm p}\Sigma)^T$ [54]. However, if one writes an equation of the same type as (63) at the scale of a cell problem (which seems to be a justified choice, because the material is isotropic at that scale), and then homogenizes, one ends up with a material for which

the Mandel stress tensor Σ no longer obeys the symmetry condition $B_{\rm p}\Sigma =$ 626 $(\boldsymbol{B}_{\mathrm{p}}\boldsymbol{\Sigma})^{T}$. This is because the material is not isotropic at the macroscale 627 and, thus, the description of remodeling based on $B_{\rm p}$ becomes inadequate. 628 Therefore, if one wants to homogenize, one should start with evolution laws 629 at the microscale, which have to be suitable to account for anisotropy, even 630 though the single constituents are isotropic at that scale. These considerations 631 lead us to Equation (52), as suggested in [22, 23], and subsequently employed 632 in [19]. 633

Remark 5. Equations (50) and (51) can be obtained by adhering to the philosophy presented in [12, 18], and subsequently adopted, for example, in [3] for growth, in [44] for growth and remodeling, and in [31, 32] for remodeling only. Accordingly, \mathbf{F}_{p} is regarded as the kinematic descriptor of the structural degrees of freedom of the medium, and $\dot{\mathbf{F}}_{p}$ as the generalized velocity with which the structural changes occur. Within this setting, it can be proven that for growth and remodeling problems, the dissipation inequality reads

$$\mathcal{D} = \boldsymbol{Y}_{\nu} : \boldsymbol{L}_{\mathrm{p}} + \mathcal{D}_{\mathrm{nc}} \ge 0, \tag{64}$$

where $\mathcal{D}_{\text{mech}} := \mathbf{Y}_{\nu} : \mathbf{L}_{\text{p}}$ is the mechanical contribution to dissipation, with \mathbf{Y}_{ν} being the dissipative part of a generalized internal force, dual to \mathbf{L}_{p} . In our work, however, \mathbf{Y}_{ν} can be identified with the tensor $\mathbf{Y}_{\nu} \equiv J_{\text{p}}^{-1} \mathbf{F}_{\text{p}}^{-\text{T}} \Sigma \mathbf{F}_{\text{p}}^{\text{T}}$, so that $\mathcal{D}_{\text{mech}}$ coincides with the mechanical dissipation encountered in the standard formulation of Elastoplasticity, i.e., $\mathcal{D}_{\text{mech}} = J_{\text{p}}^{-1} \mathbf{F}_{\text{p}}^{-\text{T}} \Sigma \mathbf{F}_{\text{p}}^{\text{T}}$: $\mathbf{L}_{\text{p}} =$ $J_{\text{p}}^{-1} \Sigma : \mathbf{F}_{\text{p}}^{-1} \dot{\mathbf{F}}_{\text{p}}$.

In the terminology of [45, 30], \mathcal{D}_{nc} is referred to as "non-compliant" contribution to the overall dissipation. Physically, it summarizes a class of phenomena that are not —or cannot be— resolved in terms of mechanical power at the scale at which the dissipation inequality is written. For instance, in the case of growth, \mathcal{D}_{nc} may represent biochemical effects contributing to the overall dissipation.

The inequality (64) can be studied in several ways, depending on the prob-653 lem at hand. First, we consider a growth problem. To this end, we assume 654 that \mathcal{D}_{nc} can be written as $\mathcal{D}_{nc} = r\mathcal{A}$, where r is the rate at which mass 655 is added or depleted from the system (its units are given by the reciprocal 656 of time), and \mathcal{A} is the energy density (per unit volume) associated with the 657 introduction or uptake of mass. In this setting, it is possible to conceive a 658 particular state of the system in which the mechanical stress is null, i.e., 659 $\Sigma = 0$, while r and A are generally nonzero. When this occurs, the system 660

⁶⁶¹ grows without mechanical dissipation, i.e., $\mathcal{D}_{mech} = 0$, whereas the overall ⁶⁶² dissipation of the system reduces to the non-compliant one:

$$\mathcal{D} \equiv \mathcal{D}_{\rm nc} = r\mathcal{A} \ge 0. \tag{65}$$

The second case addresses the situation of pure remodeling, for which we set $\mathcal{D}_{nc} = 0$, so that the dissipation inequality (64) becomes

$$\mathcal{D} = \mathcal{D}_{\text{mech}} = \boldsymbol{Y}_{\nu} : \boldsymbol{L}_{\text{p}} = J_{\text{p}}^{-1} \boldsymbol{\Sigma} : \boldsymbol{F}_{\text{p}}^{-1} \dot{\boldsymbol{F}}_{\text{p}} \ge 0.$$
(66)

It is possible to show that the evolution laws (50) and (51) are in harmony with (66).

667 6. A computational scheme for small deformations

The macro-scale model given by the problems (46) and (61), together 668 with the auxiliary cell problems (43) and (45), requires dedicated numerical 669 schemes which are subject of our current investigations. The main compu-670 tational challenge is due to the fact that the local problems depend on the 671 macro-scale in a time-dependent way. Therefore, at each time, there is a dif-672 ferent cell problem at each macroscopic point $X \in \mathcal{B}_h$. Moreover, one has to 673 transfer the information (represented by the geometry, material coefficients, 674 and unknowns of the problem) from the cell problems to the homogenized 675 problem in the domain \mathcal{B}_h , and vice versa. 676

Here, as a first step towards the numerical study of this kind of problems, we propose an algorithm adapted from [31] that could be useful in our case. In [31] it is introduced a computational algorithm, named Generalised Plasticity Algorithm (GPA), to study the mechanical response of a biological tissue that undergoes large deformations and remodeling of its internal structure. Following [31], the discrete and linearized version of the problem constituted by Equations (43), (45), (46) and (61) is formulated in three steps.

First step. The weak form of the cell problems (43) and (45), and of the homogenized problem (46) can be *formally* rewritten as

$$\mathcal{L}_1^w(\boldsymbol{\xi}, \boldsymbol{F}_p^{(0)}, \tilde{\boldsymbol{\xi}}) = 0, \qquad (67a)$$

$$\mathcal{L}_{2}^{w}(\boldsymbol{\omega}, \boldsymbol{F}_{\mathrm{p}}^{(0)}, \tilde{\boldsymbol{\omega}}) = 0, \tag{67b}$$

$$\mathcal{H}_1^w(\boldsymbol{u}^{(0)}, \boldsymbol{F}_p^{(0)}, \tilde{\boldsymbol{u}}^{(0)}) = 0, \qquad (67c)$$

where $\tilde{\boldsymbol{\xi}}$, $\tilde{\boldsymbol{\omega}}$ and $\tilde{\boldsymbol{u}}^{(0)}$ are test functions defined in certain Sobolev spaces, and \mathcal{L}_1^w , \mathcal{L}_2^w and \mathcal{H}_1^w are suitable integral operators. Together with (67a)-(67c), we rewrite in operatorial form also the homogenized problem (61) as

$$\mathcal{H}_2(\boldsymbol{\xi}, \boldsymbol{\omega}, \boldsymbol{u}^{(0)}, \boldsymbol{F}_{\mathrm{p}}^{(0)}) = \boldsymbol{0}.$$
 (68)

Note that (68) is not a weak form because the corresponding equation does not involved spatial derivatives of $F_{p}^{(0)}$.

⁶⁹¹ Second step. We perform a backward Euler method [78] for discretizing the ⁶⁹² evolution law for $F_{\rm p}^{(0)}$ given by (68), thereby ending up with the following ⁶⁹³ system of time-discrete equations,

$$\mathcal{L}_{1[n]}^{w}(\boldsymbol{\xi}_{[n]}, \boldsymbol{F}_{p[n]}^{(0)}, \tilde{\boldsymbol{\xi}}) = 0,$$
(69a)

$$\mathcal{L}_{2[n]}^{w}(\boldsymbol{\omega}_{[n]}, \boldsymbol{F}_{\mathbf{p}[n]}^{(0)}, \tilde{\boldsymbol{\omega}}) = 0, \qquad (69b)$$

$$\mathcal{H}_{1[n]}^{w}(\boldsymbol{u}_{[n]}^{(0)}, \boldsymbol{F}_{p[n]}^{(0)}, \tilde{\boldsymbol{u}}^{(0)}) = 0, \qquad (69c)$$

$$\mathcal{H}_{2[n]}(\boldsymbol{\xi}_{[n]}, \boldsymbol{\omega}_{[n]}, \boldsymbol{u}_{[n]}^{(0)}, \boldsymbol{F}_{p[n]}^{(0)}) = \mathbf{0},$$
(69d)

where n = 1, ..., N enumerates the nodes of a suitable time grid. We notice that an explicit time discrete method could be also used. However, when dealing with problems in Elastoplasticity, this election could lead to a less accurate solution.

Third step. The operators $\mathcal{L}_{1[n]}^{w}$, $\mathcal{L}_{2[n]}^{w}$, $\mathcal{H}_{1[n]}^{w}$ and $\mathcal{H}_{2[n]}$, are linear in $\boldsymbol{\xi}_{[n]}$, $\boldsymbol{\omega}_{[n]}$ and $\boldsymbol{u}_{[n]}^{(0)}$, respectively, but they are nonlinear in $\boldsymbol{F}_{p[n]}^{(0)}$. Thus, to search the solution to (69a)-(69d), we linearize at each time step according to Newton's method (with a linesearch). Therefore, at the kth iteration, $k \in \mathbb{N}, k \geq 1$, $\boldsymbol{F}_{p[n,k]}^{(0)}$ is written as

$$\boldsymbol{F}_{\mathrm{p}[n,k]}^{(0)} = \boldsymbol{F}_{\mathrm{p}[n,k-1]}^{(0)} + \boldsymbol{\Psi}_{[n,k]}, \tag{70}$$

where $F_{p[n,k-1]}^{(0)}$ is known and $\Psi_{[n,k]}$ represents the unknown increment. We introduce the notation

$$\mathcal{L}_{1[n,k-1]}^{w}(\boldsymbol{\xi}_{[n]},\tilde{\boldsymbol{\xi}}) = \mathcal{L}_{1[n]}^{w}(\boldsymbol{\xi}_{[n]},\boldsymbol{F}_{\mathrm{p}[n,k-1]}^{(0)},\tilde{\boldsymbol{\xi}}),$$
(71a)

$$\mathcal{L}_{2[n,k-1]}^{w}(\boldsymbol{\omega}_{[n]},\tilde{\boldsymbol{\omega}}) = \mathcal{L}_{2[n]}^{w}(\boldsymbol{\omega}_{[n]},\boldsymbol{F}_{\mathbf{p}[n,k-1]}^{(0)},\tilde{\boldsymbol{\omega}}),$$
(71b)

$$\mathcal{H}_{1[n,k-1]}^{w}(\boldsymbol{u}_{[n]}^{(0)}, \tilde{\boldsymbol{u}}_{[n]}^{(0)}) = \mathcal{H}_{1[n]}^{w}(\boldsymbol{u}_{[n]}^{(0)}, \boldsymbol{F}_{p[n,k-1]}^{(0)}, \tilde{\boldsymbol{u}}_{[n]}^{(0)}).$$
(71c)

 $_{705}$ Now, for each time step, and at the *k*th iteration, we solve

$$\mathcal{L}_{1[n,k-1]}^{w}(\boldsymbol{\xi}_{[n]}, \tilde{\boldsymbol{\xi}}) = 0, \qquad (72a)$$

$$\mathcal{L}_{2[n,k-1]}^{w}(\boldsymbol{\omega}_{[n]},\tilde{\boldsymbol{\omega}})=0, \qquad (72b)$$

$$\mathcal{H}_{1[n,k-1]}^{w}(\boldsymbol{u}_{[n]}^{(0)}, \tilde{\boldsymbol{u}}^{(0)}) = 0, \qquad (72c)$$

and obtain the "temporary" solutions $\boldsymbol{\xi}_{[n,k-1]}$, $\boldsymbol{\omega}_{[n,k-1]}$, and $\boldsymbol{u}_{[n,k-1]}^{(0)}$, respectively. Then, upon setting

$$\mathcal{H}_{2[n,k-1]} = \mathcal{H}_{2[n]}(\boldsymbol{\xi}_{[n,k-1]}, \boldsymbol{\omega}_{[n,k-1]}, \boldsymbol{u}_{[n,k-1]}^{(0)}, \boldsymbol{F}_{p[n,k-1]}^{(0)}),$$
(73a)

$$\mathscr{H}_{[n,k-1]} = \mathscr{H}_{[n]}(\boldsymbol{\xi}_{[n,k-1]}, \boldsymbol{\omega}_{[n,k-1]}, \boldsymbol{u}_{[n,k-1]}^{(0)}, \boldsymbol{F}_{\mathrm{p}[n,k-1]}^{(0)}),$$
(73b)

we linearize (69d), i.e.,

$$\mathcal{H}_{2[n,k-1]} + \mathscr{H}_{[n,k-1]} : \Psi_{[n,k]} = \mathbf{0}, \tag{74}$$

where $\mathscr{H}_{[n,k-1]}$ is a fourth-order tensor given by the Gâteaux derivative of $\mathcal{H}_{2[n]}$, computed with respect to its fourth argument, and evaluated in $F_{p[n,k-1]}^{(0)}$.

If the residuum $\mathbf{F}_{p[n,k]}^{(0)}$ for k greater than, or equal to, a certain k_* is less than a tolerance $\delta > 0$, then we set $\mathbf{F}_{p[n]}^{(0)} \equiv \mathbf{F}_{p[n,k_*]}^{(0)} = \mathbf{F}_{p[n,k_*-1]}^{(0)} + \Psi_{[n,k_*]}$ and we regard it as the solution of Newton's method. Thus, we compute $\boldsymbol{\xi}_{[n]}, \boldsymbol{\omega}_{[n]}$ and $\boldsymbol{u}_{[n]}^{(0)}$.

These three steps are summarized in the algorithm 1.

717 7. Numerical results

In this section, the potentiality of our model, which is given by Equations (43), (45), (46) and (61), is shown by performing numerical simulations. In particular, we make the following considerations.

(i) Geometry. We consider the composite body $\mathcal{B}^{\varepsilon}$ to have a layered threedimensional structure, and we assume that the layers are orthogonal to the direction \mathcal{E}_3 , where $\{\mathcal{E}_A\}_{A=1}^3$ is an orthonormal basis of a system of Cartesian coordinates $\{X_A\}_{A=1}^3$. In this particular case, the material properties of the heterogeneous body only change along the \mathcal{E}_3 direction and, thus, they depend solely on the coordinate X_3 . Consequently, the benchmark test at

Algorithm 1

1:	procedure				
2:	for $n = 1, \ldots, N$ do				
3:	State $k = 1$ (a)				
4:	while $e > \delta$ do (Known $F_{p[n,k-1]}^{(0)}$)				
5:	Solve $\mathcal{L}_{1[n,k-1]}^w$ and $\mathcal{L}_{2[n,k-1]}^w$ (To find $\boldsymbol{\xi}_{[n,k-1]}$ and $\boldsymbol{\omega}_{[n,k-1]}$)				
6:	Solve $\mathcal{H}^w_{1[n,k-1]}$ (To find $oldsymbol{u}^{(0)}_{[n,k-1]}$)				
7:	Solve $\mathcal{H}^w_{1[n,k-1]}$ (To find $\Psi_{[n,k]}$)				
8:	$\boldsymbol{F}_{\mathrm{p}[n,k-1]}^{(0)} \leftarrow \boldsymbol{F}_{\mathrm{p}[n,k-1]}^{(0)} + \boldsymbol{\Psi}_{[n,k]}$				
9:	Compute e				
10:	k = k + 1				
11:	end while				
12:	$oldsymbol{F}_{\mathrm{p}[n]}^{(0)} = oldsymbol{F}_{\mathrm{p}[n,k-1]}^{(0)} + oldsymbol{\Psi}_{[n,k]}$				
13:	Solve $\mathcal{L}_{1[n]}^w$ and $\mathcal{L}_{2[n]}^w$ (To find $\boldsymbol{\xi}_{[n]}$ and $\boldsymbol{\omega}_{[n]}$)				
14:	Solve $\mathcal{H}^w_{1[n]}$ (To find $oldsymbol{u}^{(0)}_{[n]}$)				
15:	Update micro and macro geometries				
16:	end for				
17: end procedure					

hand can be recast into a one dimensional problem, that is, the reference configuration of the periodic cell and the body are considered to be the unidimensional domains $\mathcal{Y}_0 = [0, \ell]$ and $\mathcal{B}_h = [0, L]$, respectively. We denote with ℓ and L, respectively, the dimension of the periodic cell and the body along the direction \mathcal{E}_3 . Moreover, we suppose that the interface Γ_0 is the middle point $\ell/2$, so that, each material under consideration has the same volume in the microscopic cell \mathcal{Y}_0 .

(ii) Material properties. We prescribe the elasticity tensor $\mathscr{C}^{\varepsilon}$ to be independent on the macroscale variable X_3 , i.e. $\mathscr{C}^{\varepsilon}(X_3) = \mathscr{C}(X_3, Y_3) \equiv \mathscr{C}(Y_3)$, where $\{Y_A\}_{A=1}^3$ is a system of microscale Cartesian coordinates. In addition, as stated above, we consider that the constituents of the heterogeneous material are isotropic, which implies that the non zero components of the 6×6 symmetric matrix representation of \mathscr{C} are given by

$$[\mathscr{C}]_{11} = [\mathscr{C}]_{22} = [\mathscr{C}]_{33} = \lambda + 2\mu, \tag{75a}$$

$$[\mathscr{C}]_{12} = [\mathscr{C}]_{13} = [\mathscr{C}]_{23} = \lambda, \tag{75b}$$

$$[\mathscr{C}]_{44} = [\mathscr{C}]_{55} = [\mathscr{C}]_{66} = \frac{1}{2}([\mathscr{C}]_{11} - [\mathscr{C}]_{12}) = \mu, \tag{75c}$$

⁷⁴⁰ where λ and μ are Lamé's parameters. We suppose that \mathscr{C} is piece-wise ⁷⁴¹ constant, which means that λ and μ are defined as

$$\lambda(Y_3) = \begin{cases} \lambda_1, & \text{in } \mathcal{Y}_0^1 \\ \lambda_2, & \text{in } \mathcal{Y}_0^2 \end{cases} \quad \text{and} \quad \mu(Y_3) = \begin{cases} \mu_1, & \text{in } \mathcal{Y}_0^1 \\ \mu_2, & \text{in } \mathcal{Y}_0^2 \end{cases}.$$
(76)

Furthermore, we consider that γ has the same value in both constituents, which means that it is already averaged.

(iii) Plastic-like distortions. We assume that the matrix representa-744 tion of the tensor $F_{\rm p}^{(0)}$ is diagonal with non-zero components $[F_{\rm p}^{(0)}]_{11} = \frac{1}{\sqrt{p}}$ 745 $[\mathbf{F}_{p}^{(0)}]_{22} = \frac{1}{\sqrt{p}}$ and $[\mathbf{F}_{p}^{(0)}]_{33} = p$, where p is defined as the remodeling pa-746 rameter. Furthermore, we restrict our investigation to the simpler case of 747 $F_{p}^{(0)}$ depending solely on X₃. This means that, the plastic-like distortions of 748 order $\varepsilon^{\bar{0}}$ are, in a sense, already averaged, and thus variable from one cell 749 to the other, not inside them. In other words, we are interested in the pro-750 duction of distortions in the tissue starting from the cell scale, rather than 751 from the cell's microstructure. This, of course, does not mean that the cell's 752 microstructure does not change. 753

Together the with assumption (ii), we find that the 6×6 matrix representation of the elasticity tensor, pulled-backed to the reference configuration, is symmetric, and its non-zero components are given by

$$[\mathscr{C}_{\mathbf{R}}]_{11} = [\mathscr{C}_{\mathbf{R}}]_{22} = (\lambda + 2\mu)p^2, \qquad [\mathscr{C}_{\mathbf{R}}]_{33} = (\lambda + 2\mu)p^{-4}, \tag{77a}$$

$$]_{12} = \lambda p^2,$$
 $[\mathscr{C}_{\mathbf{R}}]_{44} = [\mathscr{C}_{\mathbf{R}}]_{55} = \mu p^{-1},$ (77b)

$$[\mathscr{C}_{R}]_{13} = [\mathscr{C}_{R}]_{23} = \lambda p^{-1}, \qquad [\mathscr{C}_{R}]_{66} = \mu p^{2}.$$
 (77c)

⁷⁵⁷ We remark that $\mathscr{C}_{\mathbb{R}}$ depends on X_3 and time through p, whereas it inherits ⁷⁵⁸ the dependence of \mathscr{C} on the micro-scale variable, Y_3 .

 $[\mathscr{C}_{\mathrm{R}}]$

(iv) Initial and boundary conditions. In the present context, we im-759 pose Dirichlet conditions for $\boldsymbol{u}^{(0)}$ on the whole boundary $\partial \mathcal{B}_h$, i.e. we do not 760 consider a Neumann condition and therefore, $\partial_u \mathcal{B}_h \equiv \partial \mathcal{B}_h$. We note that, 761 although the homogenization process was developed for mixed boundary con-762 ditions, the whole procedure stands, since the type of boundary conditions 763 does not play a role in the derivation of the homogenized model. In par-764 ticular, we set $[\boldsymbol{u}^{(0)}]_3 = 0$ at $X_3 = 0$, and $[\boldsymbol{u}^{(0)}]_3 = \frac{u_L t}{t_f}$ at $X_3 = L$, where 765 u_L is a target value for the displacement in the direction \mathcal{E}_3 . Moreover, 766

we enforce an initial spatial distribution for the remodeling parameter p as $p_{\text{in}}(X_3) = \alpha + \beta \cos(\frac{\pi}{L}X_3)$, where α and β are constants, such that $p_{\text{in}}(X_3)$ is always strictly positive.

770 7.1. Discussion of the numerical results

Given the above considerations, we solve the following homogenized equations for $\boldsymbol{u}^{(0)}$ and p,

$$-\frac{\partial}{\partial X_3}([\hat{\mathscr{C}}_{\mathrm{R}}]_{i3n3}\frac{\partial [\boldsymbol{u}^{(0)}]_n}{\partial X_3}) = \frac{\partial [\hat{\boldsymbol{D}}_R]_{i3}}{\partial X_3}, \quad \text{for } i = 1, 2, 3,$$
(78a)

$$\langle [\boldsymbol{C}_{\rm lin}^{(0)}]_{33} \rangle \frac{\partial p}{\partial t} = \frac{\gamma}{3} \langle \operatorname{dev}(\boldsymbol{\Sigma}_{\rm lin}^{(0)}) \rangle p - \gamma \langle [\mathscr{C}_{\rm R}]_{33nn} [\boldsymbol{E}_{\rm p}]_{nn} ([\boldsymbol{C}_{\rm lin}^{(0)}]_{33} - 1) \rangle p, \quad (78b)$$

T73 The coefficients $[\hat{\mathscr{C}}_{R}]_{ijkl}$, $[\hat{D}_{R}]_{ij}$ and $[C_{lin}^{(0)}]_{ij}$ are given by Equations (47a), (47b) and (62), respectively, and are to be found by solving the auxiliary cell problems for $\boldsymbol{\xi}$ and $\boldsymbol{\omega}$, given by

$$-\frac{\partial}{\partial Y_3}([\mathscr{Q}]_{i3i3}\frac{\partial[\boldsymbol{\xi}]_{ik3}}{\partial Y_3}) = \frac{\partial[\mathscr{Q}]_{i3i3}}{\partial Y_3}\delta_{ik}, \quad \text{for } i,k=1,2,3,$$
(79a)

$$-\frac{\partial}{\partial Y_3}([\mathscr{Q}]_{i3i3}\frac{\partial[\omega]_i}{\partial Y_3}) = -\frac{\partial[\mathbf{Q}]_{33}}{\partial Y_3}\delta_{i3}, \quad \text{for } i = 1, 2, 3,$$
(79b)

776 with

$$[\mathscr{Q}]_{i3i3} = [\mathscr{C}_{\mathrm{R}}]_{i3i3} - [\mathbf{Q}]_{33}, \quad [\mathbf{Q}]_{33} = [\mathscr{C}_{\mathrm{R}}]_{33nn} [\mathbf{E}_{\mathrm{p}}]_{nn}.$$
(80a)

In this work, we are not interested to address a real world situation. Our aim is, instead, to show how the present theoretical framework can be numerically simulated. For this reason, the parameters used in our computations are arbitrarily chosen (see Table 1).

In Fig. 2, it is plotted the time evolution of the remodeling parameter 781 p at two different points of the macroscopic domain, that is at $X_3 = 7 \text{ cm}$ 782 and $X_3 = 21 \,\mathrm{cm}$. We observe that the evolution of p is quite different at 783 these two points. Indeed, at $X_3 = 21 \text{ cm}$, p increases and it is always greater 784 than one. On the contrary, at $X_3 = 7 \,\mathrm{cm}$, it is monotonically decreasing 785 and tends to be lower than one. In Fig. 3, we show the spatial profile of the 786 effective coefficients $[\mathscr{C}]_{33}$, $[\mathscr{C}_{R}]_{33}$ and $[D_{R}]_{33}$. The effective coefficient $[\mathscr{C}]_{33}$ 787 (see Remark 3) can be computed by using the analytical formula (see e.g. 788 [56, 69]),789

Parameter	Unit	Value	Parameter	Unit	Value
	[cm]	28.000 1.0000	$egin{array}{c} \lambda_1 \ \lambda_2 \end{array}$	[Pa] [Pa]	$1.00 \\ 2.00$
$u_L \ \gamma$	[1/s]	1.0000	μ_1	[Pa]	0.10
lpha eta eta	[—] [—]	$1.0035 \\ -0.0035$	$\mu_2 \ t_0$	[Pa] [s]	$\begin{array}{c} 0.06 \\ 0.00 \end{array}$
N	[—]	4.0000	t_f	$[\mathbf{s}]$	10.0

Table 1: Parameters used in the numerical simulations.

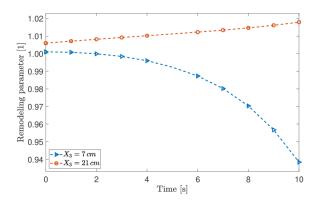


Figure 2: Evolution of the remodeling parameter p at two different points ($X_3 = 7 \text{ cm}$ and $X_3 = 21 \text{ cm}$) of the macroscopic domain.

$$\begin{aligned} [\widehat{\mathscr{C}}]_{ijkl} &= \langle [\mathscr{C}]_{ijkl} - [\mathscr{C}]_{ijp3} ([\mathscr{C}]_{p3s3})^{-1} [\mathscr{C}]_{s3kl} \rangle \\ &+ \langle [\mathscr{C}]_{ijp3} ([\mathscr{C}]_{p3s3})^{-1} \rangle \langle ([\mathscr{C}]_{s3t3})^{-1} \rangle^{-1} \langle ([\mathscr{C}]_{t3m3})^{-1} [\mathscr{C}]_{m3kl} \rangle. \end{aligned}$$
(81)

We observe that even if a loading ramp condition has been imposed on $\boldsymbol{u}^{(0)}$ 790 at the border $X_3 = L$, the effective coefficient $[\hat{\mathscr{C}}]_{33}$ does not vary on time. 791 This is because, in contrast to the case in which the plastic-like distortions 792 are accounted for, the cell and homogenized problems (cf. (48) and (49)) are 793 decoupled. On the other hand, the pulled-back effective coefficients $[\hat{\mathscr{C}}_{R}]_{33}$ 794 and $[\hat{D}_R]_{33}$, given by Equations (47a) and (47b), respectively, do change in 795 time since their equations are coupled with an evolution one and, as it can 796 be observed, they are strongly influenced by the initial distribution of p. In 797 fact, at the spatial point $X_3 = 21 \text{ cm}$, that is, when p > 1, $[\mathscr{C}_R]_{33}$ decreases 798

⁷⁹⁹ and $[\hat{D}_R]_{33}$ increases with time. The contrary occurs at $X_3 = 7$ cm, i.e. when ⁸⁰⁰ p < 1.

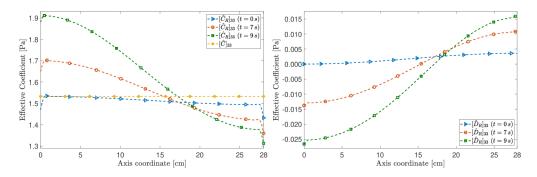


Figure 3: Spatial distribution of the effective coefficients $[\hat{\mathcal{C}}]_{33}$, $[\hat{\mathcal{C}}_{R}]_{33}$ and $[\hat{D}_{R}]_{33}$ at different time instants.

Additionally, in Fig. 4 it is illustrated the third component of the macro-801 scopic leading order term of the displacement u^{ε} at three different time 802 instants. Particularly, we plot the numerical solution of the homogenized 803 problems (46) and (49), represented with $[\boldsymbol{u}_{\mathrm{R}}^{(0)}]_3$ and $[\boldsymbol{u}^{(0)}]_3$, respectively. We 804 note that, as expected from our election of the boundary condition, the dis-805 placement component increases monotonically in time. However, we notice 806 that the introduction of the plastic-like distortions has a direct impact on the 807 displacement distribution in the interior macroscopic points. Specifically, in 808 these points the displacement has a higher magnitude. 809

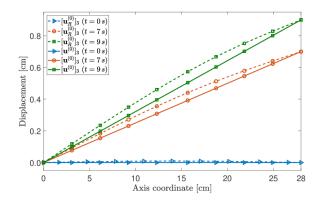


Figure 4: Spatial distribution of the macroscopic leading order term of the displacement with remodeling $([\boldsymbol{u}_{\mathrm{B}}^{(0)}]_3)$ and without remodeling $([\boldsymbol{u}^{(0)}]_3)$.

The situation described in our numerical simulations, although simplified,

could be a good starting point in the study of the remodeling of biological tissues. For example, the geometrical properties of bone's osteons permit to model them as layered composites (see e.g. [69]).

814 8. Concluding remarks

In the present work, we studied the dynamics of a heterogeneous material, constituted by two hyperlastic media with evolving micro-structure, by the application of the asymptotic homogenization technique. The evolution of the micro-structure of the composite media was characterized through the development of plastic-like distortions, which were described by means of the BKL decomposition.

The asymptotic homogenization method was applied to a set of problems 821 comprising a scale-dependent, quasi-static law of balance of linear momentum 822 and an evolution law for the tensor of plastic-like distortions. After obtaining 823 the local and homogenized problems, we rewrote them by considering the De 824 Saint-Venant strain energy density within the limit of small deformations. 825 Although the selection of the strain energy density was due to its simplicity, 826 it is helpful for the description of remodeling processes undergoing small 827 deformations. For instance, this could be the case for describing bone aging. 828 Then, the theoretical setting developed in the present work is applicable 829 (Elastoplasticity is actually quite appropriate to model the bone [73]). In 830 such a case, appropriate constitutive laws describing the progression of the 831 material properties should be found based on experimental literature (e.g. 832 [35]). Nevertheless, for studying a larger range of problems, we need to select 833 nonlinear constitutive laws and write the corresponding cell and homogenized 834 problems. 835

As a consequence of the introduction of the tensor of plastic distortions, 836 two independent cell problems were inferred, which reduce to the classical cell 837 problems encountered in the homogenization of linear problems in elastostat-838 ics. Moreover, we proposed an evolution equation for the inelastic distortions 839 describing a remodeling process. Such evolution law models a stress-driven 840 production of inelastic distortions, as the one that is often encountered in 841 studies of inelastic processes constructed on the decomposition given by (5)842 [78]. The evolution law is suitable for the case of finite strain Elastoplastic-843 ity, and for the case of remodeling of biological tissues. Finally, we outlined 844 a computational procedure in order to solve the up-scaled problems and we 845 performed numerical simulations for a particular case of a layered composite 846

⁸⁴⁷ body. Besides, we assumed that the leading order term of the asymptotic ⁸⁴⁸ expansion of the tensor of plastic distortions, $F_{\rm p}^{(0)}$, depends only on the ⁸⁴⁹ macro-scale variable X. This consideration, however, might be relaxed by ⁸⁵⁰ allowing $F_{\rm p}^{(0)}$ to take into account the heterogeneities of the composite mate-⁸⁵¹ rial through the microscopic spatial variable Y. The numerical results showed ⁸⁵² the influence of the plastic-like distortions on both the effective coefficients ⁸⁵³ and the macroscopic leading order term of the displacement.

As future work, we intend to deal with the resolution of a particular problem, like for instance the modeling of bones [49], tumor growth [67, 2, 43, 52, 70, 71], or tissue aging [20]. A further step could be the study, with the aid of the Homogenization Theory, of the coupling between the results presented in this work and the fluid flow in a hydrated tissue, or in the case of wavy laminar structures.

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⁸⁶⁹ Declaration of interest

The Authors declare that they have no conflict of interest.

871 Article information

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