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Anelastic reorganisation of fibre-reinforced biological tissues

Salvatore Di Stefano¹ · Melania Carfagna¹ · Markus M. Knodel² · Kotaybah Hashlamoun^{3,4} · Salvatore Federico⁴ · Alfio Grillo¹

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Abstract In this work, we contribute to the study of the structural reorganisation of biological tissues in response to 2 mechanical stimuli. We specialise our investigation to a class 3 of hydrated soft tissues, whose internal structure features re-4 inforcing fibres. These are oriented statistically within the 5 tissue, and their pattern of orientation is such that, at each 6 material point, the tissue is anisotropic. From its natural, stress-free state, the tissue can be distorted anelastically into 8 a global reference configuration, and then deformed under 9 the action of external mechanical loads. The anelastic dis-10 tortions are responsible for changing irreversibly the internal 11 structure of the tissue, which, in the present context, oc-12 curs through both the rearrangement of the bonds among 13 the tissue cells and the deformation-driven reorientation of 14 the fibres. The anelastic strains, in addition, are assumed to 15

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¹ Dept. of Mathematical Sciences (DISMA) "G.L. Lagrange", "*Dipartimento di Eccellenza 2018-2022*", Politecnico di Torino, C.so Duca degli Abruzzi 24, 10129, Torino (TO) Italy

E-mail: {salvatore.distefano melania.carfagna alfio.grillo}@polito.it ² Dept. of Mathematics, Chair of Applied Mathematics 1, Friedrich-Alexander-Universität Erlangen-Nürnberg, Cauerstr. 11, 91058 Erlangen, Germany

E-mail: markus.knodel@math.fau.de

³ Graduate Programme in Biomedical Engineering, The University of Calgary, 2500 University Drive NW, Calgary, AB T2N 1N4, Canada ⁴ Dept. of Mechanical and Manufacturing Engineering, The University of Calgary, 2500 University Drive NW, Calgary, AB T2N 1N4, Canada E-mail: {kwhashla salvatore.federico}@ucalgary.ca

model the onset and evolution of microcracks in the tissue, 16 which may be triggered by the mechanical loads applied to 17 the tissue in the case of traumatic events, or diseases. For 18 our purposes, we formulate an anisotropic model of remod-19 elling and we consider a fully isotropic model of structural 20 reorganisation for comparison, with the aim to study if, how, 21 and to what extent the evolution of anelastic distortions is 22 influenced by the tissue's anisotropy. 23

Keywords Anelastic distortions · Fibre-reinforcement · Biological tissues · Anisotropic media

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1 Introduction

Biological tissues tend to adapt themselves to the stimuli to 27 which they are exposed and to the environment in which 28 they are placed [60]. By "stimulus" it is meant here any 29 interaction, or combination of interactions, that yields an 30 evolution of mass, composition, shape, and internal structure 31 of a given tissue. An interaction of this kind can be genetic or 32 epigenetic, physiological or pathological, and may be related 33 to the occurrence of phenomena of various nature, associated 34 with different time and length scales. 35

In this work, emphasis is put on the evolution of the internal structure of fibre-reinforced soft tissues saturated with an interstitial fluid and exchanging mechanical interactions with it [25]. The fibres consist of collagen and are assumed to be directed according to a spatially inhomogeneous statistical distribution of orientations that makes the tissue anisotropic [42,4,30,23,18]. The interactions with the fluid are usually accounted for under the hypothesis of validity of Darcy's law [41,56,3].

Within the modelling framework outlined above, we address a type of structural reorganisation that may be associated with two types of phenomena. The first one, which is often encountered in the study of cellular aggregates and 48

Corresponding author: Alfio Grillo (alfio.grillo@polito.it)

Tel.: +39-011-0907531

Fax: +39-011-0907599

tumour spheroids, occurs through the reorganisation of the 49 extracellular matrix of the considered tissue, and leads to 50 the change of the adhesion properties of the tissue cells [55, 51 31,39]. The second phenomenon, studied in the mechanics 52 of bone, consists of the emergence of irreversible strains in 53 conjunction with the formation of micro-cracks in diseased 54 or injured tissues [27]. In spite of the fact that the aforemen-55 tioned phenomena have different nature, both of them can 56 be described by suitably re-interpreting some fundamental 57 concepts of the theory of Plasticity [46,51] (further details 58 are given in Section 2.1). More specifically, it is stipulated 59 that both the remodelling of a tissue's extra-cellular matrix 60 and the irreversible strains arising in the case of damaged or 61 overloaded tissues can be expressed in terms of plastic-like 62 distortions. The physical meaning of such distortions can be 63 captured by relating them to the concept of residual stresses, 64 which generally accompany the structural changes of a tis-65 sue. Since residual stresses persist even when all the loads 66 applied to the tissue are switched off, even an unloaded con-67 figuration, taken as reference for the tissue's evolution, may 68 happen to be in a stressed state. Accordingly, it is possible to 69 identify the plastic distortions with the transformations that 70 bring the tissue from the stressed state associated with the 71 chosen reference configuration to a stress-free state, i.e., a 72 state reached by eliminating all applied loads and relaxing 73 all residual stresses [51] (we recall that a similar definition 74 is given in [58] for the remodelling associated with growth). 75

Determining physically sound evolution laws for the dis-76 tortions characterising the structural adaptation of biological 77 tissues is a crucial task, which has been undertaken by sev-78 eral authors (see e.g. [16,45,28,1,43,29,53,36]). One of the 79 main challenges of mathematical modelling is to predict how 80 the structural evolution of a tissue is modulated by mechani-81 cal stress. This issue is particularly relevant when also other 82 phenomena, such as growth [57], mechano-transduction [6], 83 and interactions with other stimuli [44,48,10], have to be 84 accounted for. Moreover, since the formulation of models for 85 the structural evolution of tissues allows for a certain free-86 dom, and since a model that is reliable for a certain tissue 87 may be inaccurate for another one, it is difficult to find a 88 unified criterion for determining a priori how such models 89 should be constructed. To our knowledge, however, Epstein 90 and Maugin [16] prescribed a series of conditions that should 91 be satisfied in order to formulate acceptable structural evolu-92 tions. These rules, in turn, are based on the theory developed, 93 for example, in [15, 50, 14]. 94

With the purpose of seeking for a unified form of the structural evolution laws of biological tissues, we take a phenomenological law of remodelling in isotropic media [31] and, by following the rules put forward in [16], we rephrase it for the case of an anisotropic tissue. To this end, we elaborate the anisotropic hyperelastic model of fibre-reinforced tissues developed in [24, 19, 61], in which the interaction 112

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with an interstitial fluid is considered, and we extend it to 102 the case of nonlinear elastoplastic material behaviour. Then, 103 after specifying the equations governing the deformation of 104 the tissue, the fluid flow, and the evolution of the plastic-like 105 distortions, we test our model by solving numerically ded-106 icated benchmark problems. The main result of our work 107 is the evaluation of the interplay between remodelling and 108 the anisotropy of the tissue. This interplay is highlighted by 109 comparing the results of our anisotropic model with those 110 predicted by an isotropic model taken as reference [39]. 111

2 Theorethical background

We adopt with slight variations the covariant formalism of Continuum Mechanics presented in [47].

The motion of the solid phase is described in terms of a one-parameter family of embeddings $\chi_t : \mathcal{B} \to \mathcal{S}$, where $t \in \mathcal{I}$ is time and $\mathcal{I} \subseteq \mathbb{R}$ is an interval, \mathcal{S} is the threedimensional Euclidean space, and the open set $\mathcal{B} \subset \mathcal{S}$ is said to be the reference configuration of the tissue. It holds that $\chi_t(X) = \chi(X, t)$, with $\chi : \mathcal{B} \times \mathcal{I} \to \mathcal{S}$. 120

For every $x \in S$ and $X \in B$, T_xS and T_XB are the 121 tangent spaces of S and \mathcal{B} at x and X, respectively. The 122 disjoint unions $TS := \sqcup_{x \in S} T_x S$ and $TB := \sqcup_{X \in B} T_X B$ are 123 the tangent bundles of S and \mathcal{B} . The spaces dual to $T_x S$ and 124 $T_X \mathcal{B}$ are referred to as co-tangent spaces and denoted by $T_x^{\star} \mathcal{S}$ 125 and $T_X^{\star}\mathcal{B}$, while $T^{\star}\mathcal{S} := \sqcup_{x \in \mathcal{S}} T_x^{\star}\mathcal{S}$ and $T^{\star}\mathcal{B} := \sqcup_{x \in \mathcal{B}} T_x^{\star}\mathcal{B}$ 126 are the co-tangent bundles. Finally, $\mathcal B$ and $\mathcal S$ are equipped 127 with the metric tensors G and g, respectively. 128

The tangent map of $\chi_t : \mathcal{B} \to \mathcal{S}$ is the deformation gradi-129 ent tensor $F(X, t): T_X \mathcal{B} \to T_{\chi_t(X)} \mathcal{S}$. Once two local systems 130 of coordinates are chosen in \mathcal{B} and \mathcal{S} , the components of 131 **F** read $F_A^a = \partial \chi^a / \partial X^A \equiv \chi^a_{,A}$, with a, A = 1, 2, 3. The 132 determinant $J = \det F$, called *volumetric ratio*, is required 133 to be strictly positive at all points $X \in \mathcal{B}$ and at all times. 134 Another measure of deformation that will be used in the fol-135 lowing is the right Cauchy-Green deformation tensor defined 136 by $C = F^{\mathrm{T}}gF \equiv F^{\mathrm{T}}.F$. 137

2.1 Anelastic distortions and natural state

As anticipated in Section 1, the structural evolution of a tis-139 sue is interpreted as a sequence of plastic-like distortions, de-140 scribed by a *distortion tensor*. This tensor, denoted by $F_{\rm p}$, is 141 introduced by invoking the Bilby-Kröner-Lee (BKL) decom-142 position of the deformation gradient tensor. Consequently, F 143 is written as $F = F_e F_p$, where F_e is said to be the tensor 144 of elastic distortions and $J = J_e J_p$, with $J_e := det F_e > 0$ 145 and $J_p := \det F_p > 0$. In the literature, decompositions of the 146 deformation gradient tensor have been extensively used to 147 address problems of biomechanical interest (see e.g. [58, 16, 148

45, 1, 2, 26, 36, 39, 40]). The physical and geometrical mean-149 ing of the BKL decomposition have been explained in detail, 150 for instance, in [51,32] and they have been recently used to 151 study the structural evolution of a growing tumour in [10]. 152 For every pair $(X, t) \in \mathcal{B} \times \mathcal{I}$, $F_p(X, t)$ maps $T_X \mathcal{B}$ into a 153 vector space, denoted by $\mathcal{N}_X(t)$ and consisting in the im-154 age of $T_X \mathcal{B}$ through $F_p(X, t)$ [10], whose vectors represent 155 body elements in a stress-free state [11]. In general, F_{p} is 156 an incompatible tensor (see [51,59] and references therein). 157 The way in which $F_{p}(X, t)$ operates on $T_X \mathcal{B}$ is illustrated in 158 Figure 1. 159



collection of natural states

Fig. 1 Multiplicative decomposition of the deformation gradient tensor F. When the BKL decomposition is enforced, the rule $F = F_e F_p$ applies. When, instead, the decomposition \dot{a} la Epstein and Maugin [15,50] is employed, the rule $F\Pi = F_e$ is used.

In light of the BKL decomposition, each vector in the 160 natural state $\mathfrak{u}_X(t) \in \mathfrak{N}_X(t)$ can be distorted elastically into 161 $\boldsymbol{u}_{X}(t) = \boldsymbol{F}_{e}(X, t)\boldsymbol{u}_{X}(t) \in T_{X}\mathcal{S}$, with $x = \chi_{t}(X)$. Moreover, we 162 introduce the tensor $\Pi(X, t) : \mathcal{N}_X(t) \to T_X \mathcal{B}$ as the inverse of 163 $F_{p}(X, t)$, so that the relation $U_{X} = \Pi(X, t)\mathfrak{u}_{X}(t) \in T_{X}\mathcal{B}$ holds 164 true. Finally, we notice that, since $F(X, t) : T_X \mathcal{B} \to T_X \mathcal{S}$ is 165 such that $u_x(t) = F(X, t)U_X$, with $x = \chi_t(X)$, it also holds 166 true that 167

$$\boldsymbol{u}_{\boldsymbol{X}}(t) = \boldsymbol{F}(\boldsymbol{X}, t)\boldsymbol{U}_{\boldsymbol{X}} = \boldsymbol{F}(\boldsymbol{X}, t)\boldsymbol{\Pi}(\boldsymbol{X}, t)\boldsymbol{\mathfrak{u}}_{\boldsymbol{X}}(t)$$
$$= \boldsymbol{F}_{\mathrm{e}}(\boldsymbol{X}, t)\boldsymbol{\mathfrak{u}}_{\boldsymbol{X}}(t). \tag{1}$$

It follows thus from this chain of equalities, which has to be 168 respected for all $\mathfrak{u}_X(t) \in \mathfrak{N}_X(t)$, that the elastic distortion 169 tensor is given by $F_e = F \Pi$. We remark that this result goes 170 far behind the simple renaming of F_{p}^{-1} with Π , for it actually 171 discloses the possibility of exploring some analogies of the 172 BKL decomposition with the theory of material uniformity 173 [15,50,16,14,54] (quoting verbatim from [16] "a body is 174 said to be materially uniform if all its points are made of the 175 176 same material"). However, we do not speculate here on this analogy because it is out of the scope of our work. 177

2.2 The fibre pattern

Following the framework presented in [24, 19, 61, 5, 34, 35], 179 also in this work we study fibre-reinforced tissues in which 180 the fibres are oriented statistically. The first assumption of 181 our approach is that, at each material point X that finds it-182 self in a natural state, the tissue is transversely isotropic with 183 respect to the direction associated with the unit vector $\mathbf{m}_{\mathbf{X}}$, 184 which defines the direction of local alignment of the fibre 185 passing through X. The second assumption is that the fibres' 186 directional distribution is such that the tissue as a whole 187 is transversely isotropic with respect to a global symmetry 188 axis, identified with the unit vector \mathbf{m}_0 . Moreover, in the 189 sequel we restrict our attention to a sample of tissue charac-190 terised by cylindrical shape and material properties that vary 191 only along its geometrical axis. The sample is thus homoge-192 neous on each cross section. A consequence of this setting is 193 that the sample's geometrical axis coincides with the axis of 194 transverse isotropy, which is also then symmetry axis of the 195 tissue. 196

To account for the statistical orientation of the fibres, we adhere to the framework discussed in [19] and we introduce the function $\wp_X : \mathbb{S}^2 \mathcal{N}_X(t) \to \mathbb{R}^+_0$, with

$$\mathbb{S}^2 \mathcal{N}_X(t) := \{ \mathfrak{m}_X \in \mathcal{N}_X(t) \colon \|\mathfrak{m}_X\| = 1 \},$$
(2)

and $\wp_X(\mathfrak{m}_X)$ measuring the probability density that a (rectified) fibre passing through X be directed along \mathfrak{m}_X .

With respect to an orthonormal vector basis $\{\mathbf{e}_{\alpha}\}_{\alpha=1}^{3}$ of $\mathcal{N}_{X}(t)$, such that \mathbf{e}_{3} is parallel to \mathbf{m}_{0} , a unit vector $\mathbf{m}_{X} \in \mathcal{N}_{X}(t)$ can be expressed in spherical coordinates as $\mathbf{m}_{X} = \check{\mathbf{m}}_{X}(\vartheta, \varphi)$, where the vector-valued function $\check{\mathbf{m}}_{X}$: $[0, \pi] \times [0, 2\pi[\rightarrow 205] \mathbb{S}^{2}\mathcal{N}_{X}(t)$ is given by 206

$$\check{\mathbf{m}}_X(\vartheta,\varphi) = \sin\vartheta\cos\varphi\,\mathbf{e}_1 + \sin\vartheta\sin\varphi\,\mathbf{e}_2 + \cos\vartheta\,\mathbf{e}_3. \tag{3}$$

Accordingly, a physical quantity \mathfrak{F}_X depending on the lo-207 cal direction of fibre alignment, and thus defined over the 208 set $\mathbb{S}^2 \mathcal{N}_X(t)$, can be rewritten as a function of ϑ and φ , i.e., 209 $\mathfrak{F}_X(\mathfrak{m}_X) = \mathfrak{F}_X(\mathfrak{m}_X(\vartheta,\varphi)) = \mathfrak{F}_X(\vartheta,\varphi)$. In particular, the prob-210 ability density becomes $\wp_X(\mathfrak{m}_X) = \check{\wp}_X(\vartheta, \varphi)$ and, since the 211 tissue as a whole is assumed to be transversely isotropic with 212 respect to \mathbf{m}_0 , \check{p} is not allowed to depend on the longitude, 213 φ . Consequently, the equality $\varphi_X(\mathfrak{m}_X) = \check{\varphi}_X(\vartheta)$ must be 214 fulfilled. 215

Following the formalism adopted in [20], the directional $_{216}$ average of \mathfrak{F}_X is defined as $_{217}$

$$\begin{split} \langle\!\langle \mathfrak{F}_X(\mathfrak{m}_X) \rangle\!\rangle &= \int_{\mathbb{S}^2 \mathcal{N}_X(t)} \mathfrak{F}_X(\mathfrak{m}_X) \wp_X(\mathfrak{m}_X) \\ &= \int_0^{2\pi} \int_0^{\pi} \check{\mathfrak{F}}_X(\vartheta, \varphi) \check{\wp}_X(\vartheta) \sin \vartheta d\vartheta d\varphi. \end{split}$$
(4)

All physical quantities featuring in the mathematical model, ²¹⁸ including the probability density, are assumed to be invariant ²¹⁹

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under the reflection $\mathfrak{m}_X \mapsto -\mathfrak{m}_X$, for all \mathfrak{m}_X . This permits 220 to rephrase the directional average (4) as 221

$$\begin{split} \langle\!\!\langle \mathfrak{F}_X(\mathfrak{m}_X) \rangle\!\!\rangle &= 2 \int_{\mathbb{S}^{2+} \mathcal{N}_X(t)} \mathfrak{F}_X(\mathfrak{m}_X) \varphi_X(\mathfrak{m}_X) \\ &= \int_0^{2\pi} \!\!\!\int_0^{\pi/2} \check{\mathfrak{F}}_X(\vartheta, \varphi) \check{\psi}_X(\vartheta) \sin \vartheta d\vartheta d\varphi, \end{split}$$
(5)

where $\mathbb{S}^{2+}\mathcal{N}_X(t)$ is the "northern" hemisphere [5], i.e.,

$$\mathbb{S}^{2+}\mathcal{N}_X(t) := \{\mathfrak{m}_X \in \mathbb{S}^2\mathcal{N}_X(t) : \mathfrak{m}_X.\mathfrak{m}_0 \ge 0\},\tag{6}$$

and the probability density $\check{\psi}_X : [0, \pi/2] \to \mathbb{R}^+_0$ is defined 223 by the equality $\check{\psi}_X(\vartheta) = \psi_X(\check{\mathfrak{m}}_X(\vartheta,\varphi))$, for all $(\vartheta,\varphi) \in$ 224 $[0, \pi/2] \times [0, 2\pi[$, with $\psi_X = 2\wp_X|_{\mathbb{S}^{2+}_X \mathcal{B}}$ [5]. As done in previ-225 ous works [21,5], we assume that $\check{\psi}_X$ is the pseudo-Gaussian 226 distribution 227

$$\check{\psi}_X(\vartheta) = \frac{\check{\gamma}_X(\vartheta)}{2\pi \int_0^{\pi/2} \check{\gamma}_X(\vartheta') \sin \vartheta' d\vartheta'},\tag{7a}$$

$$\check{\gamma}_X(\vartheta) = \exp\left(-\frac{[\vartheta - \mathfrak{q}]^2}{2\omega^2}\right),$$
(7b)

where q and ω are referred to as fibre mean angle and stan-228 dard deviation, respectively. Since, as anticipated above, q 229 and ω are hypothesised to vary only along the axis of the 230 sample, they can be written as functions of the normalised 231 axial variable $\xi \in [0, 1]$, which is zero at the sample's lower 232 boundary and equal to one at the upper boudary. Hereafter, 233 we prescribe the expressions 234

$$q(\xi) = \frac{\pi}{2} \left\{ 1 - \cos\left(\frac{\pi}{2} \left[-\frac{2}{3}\xi^2 + \frac{5}{3}\xi \right] \right) \right\},\tag{8a}$$

$$\omega(\xi) = 10^3 [(1 - \xi)\xi]^4 + 3 \cdot 10^{-2}, \tag{8b}$$

which qualitatively reproduce the alignment of fibres in ar-235 ticular cartilage [52]. According to (8a) and (8b), the mean 236 angle takes on the values q(0) = 0 and $q(1) = \pi/2$, and the 237 standard deviation attains its minimum at $\xi = 0$ and $\xi = 1$. 238 Hence, the fibres are more likely to be found aligned with 239 the sample's symmetry axis at the bottom of the sample, and 240 more likely to be lying on transverse plane at the top. More-241 over, at $\xi = 1/2$, the standard deviation reaches its maximum, 242 thereby tending to randomise the fibre orientation and, con-243 sequently, to make the tissue isotropic in the middle of the 244 sample. 245

Note that $\mathbf{m} : \mathcal{B} \to \mathcal{N}(t)$ indicates the vector field such 246 that $\mathfrak{m}(X) = \mathfrak{m}_X$, and $\mathcal{N}(t) := \sqcup_{X \in \mathcal{B}} \mathcal{N}_X(t)$ is the bundle of 247 all spaces $\mathcal{N}_X(t)$. 248

2.3 Constitutive laws

At each material point, the solid phase of the tissue is mod-250 elled as a hyperelastic material. This hypothesis allows to 251 describe the mechanical behaviour of the solid phase en-252 tirely in terms of a strain energy density, and to express the 253 latter as a function of the elastic part of the deformation, 254 only. More precisely, if $W_{\rm R} = \hat{W}_{\rm R}(C, X, t)$ denotes the strain 255 energy density of the solid phase, written per unit volume 256 of the reference configuration (note that the the material in-257 homogeneities and their evolution are accounted for by the 258 explicit dependence of \hat{W}_{R} on the material points and time, 259 respectively), it is possible to write [7,16] 260

$$\hat{W}_{\rm R}(C, X, t) = \frac{1}{J_{\Pi}(X, t)} \hat{W}_{\nu}(C_{\rm e}(X, t), X), \tag{9}$$

where \hat{W}_{ν} is measured per unit volume of the natural state, 261 $J_{\Pi} = \det \Pi$ and $C_{e} = F_{e}^{T} F_{e} = F_{e}^{T} g F_{e} = \Pi^{T} C \Pi$ is the 262 elastic part of the right Cauchy-Green deformation tensor. 263 We remark that, in Equation (9), the explicit dependence 264 of the strain energy function on material points is given 265 through ξ . In the following, however, for the sake of a lighter 266 notation, the explicit dependence of \hat{W}_{ν} on material points, 267 X, is omitted but understood, and we adapt to the present 268 framework a strain energy density used in previous works 269 [24, 19, 61, 5, 40, 35], i.e., 270

$$\hat{W}_{\nu}(C_{\rm e}) = \Phi_{\rm s\nu}\hat{U}(J_{\rm e}) + \Phi_{0\rm s\nu}\hat{W}_0(C_{\rm e}) + \Phi_{1\rm s\nu}\hat{W}_{\rm en}(C_{\rm e}), \qquad (10)$$

where

$$\Phi_{0s\nu} = J_e \phi_{0s},\tag{11a}$$

$$\Phi_{1s\nu} = J_e \phi_{1s},\tag{11b}$$

$$\Phi_{\mathrm{s}\nu} = \Phi_{0\mathrm{s}\nu} + \Phi_{1\mathrm{s}\nu} = J_{\mathrm{e}}\phi_{\mathrm{s}} \tag{11c}$$

are the volumetric fractions of the non-fibrous matrix, fibres, 272 and solid phase as a whole, respectively, all measured per 273 unit volume of the natural state, while $\hat{U}(J_e)$, $\hat{W}_0(C_e)$, and 274 $\hat{W}_{en}(C_e)$ are given by 275

$$\hat{U}(J_{\rm e}) = \alpha_0 \mathcal{H}(J_{\rm cr} - J_{\rm e}) \frac{[J_{\rm e} - J_{\rm cr}]^{2q}}{[J_{\rm e} - \Phi_{\rm sv}]^r},$$
(12a)

$$\hat{W}_0(C_e) = \alpha_0 \left[\frac{\exp\left(\alpha_1 [I_{1e} - 3] + \alpha_2 [I_{2e} - 3]\right)}{[I_{3e}]^{\alpha_3}} - 1 \right], \quad (12b)$$

$$\hat{W}_{en}(\boldsymbol{C}_{e}) = \hat{W}_{1i}(\boldsymbol{C}_{e}) + \langle\!\langle \hat{W}_{1a}(\boldsymbol{C}_{e}, \boldsymbol{\mathfrak{m}}) \rangle\!\rangle.$$
(12c)

In (12a)–(12c), $\alpha_0 = 0.125$ MPa, $\alpha_1 = 0.778$, $\alpha_2 = 0.111$, 276 $\alpha_3 = \alpha_1 + 2\alpha_2 = 1, q \ge 0, \text{ and } r \in [0, 1]$ are material 277 parameters, $J_{cr} \in]\Phi_{s\nu}, 1]$ is a critical value of J_e (in this work, 278 we take q = 2, r = 0.5, and $J_{cr} = \Phi_{s\nu} + 0.1$, $I_{1e} = tr(C_e)$, 279 $I_{2e} = \frac{1}{2} \{ [tr(C_e)]^2 - tr(C_e^2) \}, \text{ and } I_{3e} = \det C_e \text{ are the principal} \}$ 280 invariants of $C_{\rm e}$, \hat{W}_{1i} is the isotropic part of the strain energy 281 density of the fibres (it has the same functional form as (12b), 282 but it features different coefficients), and $\hat{W}_{1a}(C_e, \mathfrak{m})$ reads 283

$$\hat{W}_{1a}(C_{\rm e},\mathfrak{m}) = \mathcal{H}(I_{4\rm e}-1)\frac{1}{2}c[I_{4\rm e}-1]^2,$$
(13)

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where $I_{4e} = C_e : \mathfrak{m} \otimes \mathfrak{m} = C : (\Pi \mathfrak{m} \otimes \Pi \mathfrak{m})$ and c = 7.46 MPa. In (12a) and (13), \mathcal{H} is the Heaviside function, i.e., $\mathcal{H}(s) = 1$ for all $s \ge 1$, and $\mathcal{H}(s) = 0$ for all s < 0. Finally, it is possible to define the unit vector field

$$M = \frac{\Pi \mathfrak{m}}{\|\Pi \mathfrak{m}\|}.$$
 (14)

²⁸⁸ Consequently, the structure tensor field associated with the ²⁸⁹ natural state, i.e., $a = m \otimes m$, transforms as

$$A = M \otimes M = \frac{\Pi \mathfrak{a} \Pi^{\mathrm{T}}}{(\Pi^{\mathrm{T}} \cdot \Pi) : \mathfrak{a}},$$
(15)

with *A* being the structure tensor field associated with the reference configuration, and the invariant I_{4e} becomes $I_{4e} =$

²⁹² $I_4 I_{4\Pi}$, where we used the notation

$$I_4 = \boldsymbol{C} : \boldsymbol{A},\tag{16a}$$

$$I_{4\Pi} = (\mathbf{\Pi}^{\mathrm{T}}.\mathbf{\Pi}) : \mathfrak{a}. \tag{16b}$$

²⁹³ The energy $\hat{U}(J_e)$ is zero for J_e above the critical volume ratio ²⁹⁴ J_{cr} (which, in general, is a function of material points), and ²⁹⁵ diverges for J_e tending to Φ_{sv} from above, thereby preventing ²⁹⁶ the elastic distortions from violating the unilateral constraint ²⁹⁷ $J_e \ge \Phi_{sv}$. The constitutive part of the first Piola-Kirchhoff ²⁹⁸ stress tensor associated with the solid phase is given by

$$\boldsymbol{P} = \boldsymbol{F} \left[\frac{1}{J_{\Pi}} \boldsymbol{\Pi} \left(2 \frac{\partial \hat{W}_{\nu}}{\partial \boldsymbol{C}_{\mathrm{e}}} (\boldsymbol{C}_{\mathrm{e}}) \right) \boldsymbol{\Pi}^{\mathrm{T}} \right].$$
(17)

²⁹⁹ Consequently, *P* can be expressed constitutively as a function ³⁰⁰ of *F* and Π , i.e., $P = \hat{P}(F, \Pi)$. Also in this case, the explicit ³⁰¹ dependence on material points is omitted but understood.

The system under study, comprising a porous solid phase (i.e., matrix and reinforcing fibres) and an interstitial fluid, is assumed to be saturated, thereby meaning that the porosity of the medium coincides with the volumetric fraction of the fluid, which is thus given by $\phi_f = 1 - \phi_s$.

The mathematical model presented in the following is based on the hypothesis that the interstitial fluid obeys Darcy's law. This requires the introduction of a permeability tensor for the tissue. In this work, we adapt to our problem the constitutive framework developed in [23,22,19,61,34]. Hence, we assume that the spatial permeability tensor reads [61]

$$\boldsymbol{k} = k_0 \frac{[J J_{\Pi} - \Phi_{1s\nu}]^2}{J^2 J_{\Pi}^2} \boldsymbol{g}^{-1} + k_0 \frac{[J J_{\Pi} - \Phi_{1s\nu}] \Phi_{1s\nu}}{J^2 J_{\Pi}^2} \boldsymbol{F} \boldsymbol{\Pi} \left\langle \frac{\boldsymbol{\mathfrak{a}}}{I_{4e}} \right\rangle \boldsymbol{\Pi}^{\mathrm{T}} \boldsymbol{F}^{\mathrm{T}},$$
(18)

where k_0 is taken to be of the Holmes&Mow type [41], i.e.,

$$k_0 = k_{0\nu} \left[\frac{JJ_{\Pi} - \Phi_{s\nu}}{1 - \Phi_{s\nu}} \right]^{\kappa_0} \exp\left(\frac{1}{2}m_0[J^2 J_{\Pi}^2 - 1]\right),$$
(19)

where $\kappa_0 = 0.0848$ and $m_0 = 4.638$ are model parameters, and $k_{0\nu}$ is a reference permeability. As done elsewhere (e.g. in [61]), $k_{0\nu}$ is taken as a function of the axial coordinate, ξ , and its functional form is defined in (22). From (18) and (19) we notice that, since the product $JJ_{\Pi} = J_e$ has to be greater than, or equal to, $\Phi_{s\nu}$, the permeability tensor is positive semi-definite for $JJ_{\Pi} \ge \Phi_{s\nu} \ge \Phi_{1s\nu}$ and, in particular, it is positive definite when the strict inequality is satisfied, i.e., when $JJ_{\Pi} > \Phi_{s\nu}$.

For future use, we compute the Piola transform of k, i.e., ³²³ $K = JF^{-1}kF^{-T}$, which reads ³²⁴

$$\boldsymbol{K} = k_0 \frac{[JJ_{\Pi} - \Phi_{1s\nu}]^2}{JJ_{\Pi}^2} \boldsymbol{C}^{-1} + k_0 \frac{[JJ_{\Pi} - \Phi_{1s\nu}]\Phi_{1s\nu}}{JJ_{\Pi}^2} \boldsymbol{\Pi} \left(\frac{\mathfrak{a}}{I_{4e}}\right) \boldsymbol{\Pi}^{\mathrm{T}}.$$
 (20)

Clearly, since k is positive semi-definite, K is positive semi-325 definite too. Note also that *K* can be written as $K = \hat{K}(F, \Pi)$, 326 where the dependence on F is through C because of objec-327 tivity, and the dependence on X is understood. In fact, in the 328 case of inhomogeneous materials, the dependence of $k_{0\nu}$ on 329 material points can be taken into account by expressing $k_{0\nu}$ 330 as a function of the void ratio associated with the natural 331 state, $e_{\nu} = (1 - \Phi_{s\nu})/\Phi_{s\nu}$, and specifying how the volumetric 332 fraction $\Phi_{s\nu}$ depends on the normalised axial coordinate ξ 333 (we recall, indeed, that the material is assumed here to be 334 inhomogeneous only axially). In this work, we assign the 335 volumetric fractions of matrix and fibres in the tissue's natu-336 ral state, $\Phi_{0s\nu}$ and $\Phi_{1s\nu}$, and we compute thus the volumetric 337 fraction of the solid phase as $\Phi_{s\nu} = \Phi_{0s\nu} + \Phi_{1s\nu}$. In particular, 338 we prescribe [61] 339

$$\Phi_{0s\nu} = \hat{\Phi}_{0s\nu}(\xi) = -0.062\xi^2 + 0.038\xi + 0.046, \tag{21a}$$

$$\Phi_{1s\nu} = \hat{\Phi}_{1s\nu}(\xi) = +0.062\xi^2 - 0.138\xi + 0.204, \tag{21b}$$

$$\Phi_{s\nu} = \Phi_{s\nu}(\xi) = -0.100\xi + 0.250.$$
(21c)

Following the constitutive framework adopted in previous works, we assume that $k_{0\nu}$ depends on e_{ν} as suggested by Holmes&Mow [41]. Hence, given the constant referential void ratio $e_{\nu}^{(0)} = 4$ and the constant referential scalar permeability $k_{0\nu}^{(0)} = 3.7729 \cdot 10^{-3} \text{ mm}^4(\text{Ns})^{-1}$, we assign $k_{0\nu}$ through the expression [61]

$$\frac{k_{0\nu}}{k_{0\nu}^{(0)}} = \left[\frac{e_{\nu}}{e_{\nu}^{(0)}}\right]^{\kappa_0} \exp\left(\frac{m_0}{2}\left[\left(\frac{1+e_{\nu}}{1+e_{\nu}^{(0)}}\right)^2 - 1\right]\right).$$
(22)

In summary, the constitutive framework adopted in this 346 work describes a hydrated, fibre-reinforced tissue, whose 347 solid phase is hyperelastic, transversely isotropic with respect 348 to a global symmetry axis (the direction of which is identi-349 fied by the unit vector \mathbf{m}_0), and inhomogeneous along this 350 axis. We emphasise that, within the employed approach, the 351 inhomogeneity is due to the fact that the volumetric fractions 352 of matrix and fibres, $\Phi_{0s\nu}$ and $\Phi_{1s\nu}$, the standard deviation of 353

the probability density, ω , and the mean angle of fibre orien-354 tation, q, depend on the normalised axial coordinate through 355 the expressions (8a) and (8b), which are in qualitative agree-356 ment with the histological features of articular cartilage, as 357 revealed by X-ray diffraction experiments [52]. 358

3 Description of Remodelling 359

The mathematical model of the physical system under study 360 is characterised by two dissipative phenomena. First we con-361 sider the one related to the fluid flow, which is affected by 362 dissipative forces exchanged between the fluid and the solid 363 phase. We prescribe that these forces depend linearly on 364 the filtration velocity $q = \phi_f [v_f - v_s]$ and, by disregard-365 ing the influence of gravity on the flow, we obtain Darcy's 366 law, which reads $q = -k \operatorname{grad} p$ in spatial form [3], and 367 $Q = -K \operatorname{Grad} p$ in the so-called "material" form. Here, 368 $Q := JF^{-1}q$ is the Piola transform of the filtration velocity, 369 and Grad $p = F^{T}$ grad p is the "material" pressure gradient, 370 obtained by differentiating p with respect to the coordinates 371 associated with the reference configuration. We remark that 372 the filtration velocity represents the specific mass flux vector 373 associated with the motion of the fluid relative to the solid. 374

The second dissipative phenomenon addressed in this 375 work is due to the reorganisation of the tissue's internal 376 structure. This process is described here in analogy with the 377 theory of finite strain plasticity through Π [16]. The rate with 378 which the anelastic distortions associated with Π evolve in 379 time is given by $\Lambda = \dot{\Pi} \Pi^{-1}$ and it will be referred to as *tensor* 380 of rate of remodelling. In the sequel, we shall assume that 381 remodelling is a volume-preserving process, which yields the 382 restriction $J_{\Pi} = 1$ and implies that Λ is a deviatoric second-383 order tensor. Within this framework, the generalised force 384 power-conjugate to Λ is the Mandel stress tensor $\Sigma = CS$ 385 [15,50], where $S = F^{-1}P$ is the constitutive part of the 386 second Piola-Kirchhoff stress tensor of the solid phase. 387

3.1 Dissipation Inequality 388

By accounting for the contributions due to the flow and re-389 modelling, denoted by \mathfrak{D}_{flow} and \mathfrak{D}_{rem} , respectively, the dis-390 sipation of the system under study can be written as [39] 391

$$\mathfrak{D}_{\mathsf{R}} = \underbrace{\mathbf{K} : [\operatorname{Grad} p \otimes \operatorname{Grad} p]}_{\mathfrak{D}_{\operatorname{frow}} \ge 0} \underbrace{-\boldsymbol{\Sigma} : \boldsymbol{\Lambda}}_{\mathfrak{D}_{\operatorname{rem}}} \ge 0.$$
(23)

Since the positive semi-definiteness of *K* guarantees that 392 \mathfrak{D}_{flow} is non-negative for all pressure gradients, the fulfilment 393 of the inequality $\mathfrak{D}_R \ge 0$ is equivalent to requiring the con-394 dition \mathfrak{D}_{rem} = $-\Sigma$: Λ \geq 0 for all Σ and $\Lambda.$ Moreover, the 395 physical observation that remodelling is triggered by stress 396

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398 by exploiting the fact that Σ complies, by construction, with 399 the symmetry condition $\Sigma C = CSC = (\Sigma C)^{T} [15, 50]$. Upon 400 setting Y := CSC, this yields the chain of equalities 401

$$\Sigma : \Lambda = (CSC) : (\Lambda C^{-1}) = Y : \operatorname{sym}(\Lambda C^{-1}),$$
(24)

which allows to rephrase \mathfrak{D}_{rem} as [15]

$$\mathfrak{D}_{\text{rem}} = -Y : \operatorname{sym}(\Lambda C^{-1}) \ge 0.$$
(25)

We recall that the stress tensor Y can be obtained by express-403 ing the strain energy density as a function of the Piola strain $\mathcal{E} = \frac{1}{2} [\mathbf{G}^{-1} - \mathbf{C}^{-1}] [15, 50].$

We prescribe here that **Y** and sym(ΛC^{-1}) are related to 406 each other through an expression of the type 407

$$\operatorname{sym}(\mathbf{\Lambda} \boldsymbol{C}^{-1}) = -\boldsymbol{\mathcal{R}},\tag{26}$$

where \mathcal{R} is a tensor-valued function that has to be specified 408 constitutively. Equation (26) shall also be referred to as the 409 remodelling law. 410

To satisfy the condition $\mathfrak{D}_{rem} \geq 0$, we assume here that 411 \mathcal{R} can be written as $\mathcal{R} = \mathbb{T} : Y$, where \mathbb{T} is a fourth-order 412 tensor endowed with the major symmetry and such that the 413 inequality $\mathfrak{D}_{rem} = Y : \mathbb{T} : Y \ge 0$ is respected for all Y 414 (i.e., \mathbb{T} has to be positive semi-definite). The constitutive 415 expression defining \mathbb{T} specifies the law of remodelling that 416 one is interested in. It should be noticed, however, that since 417 Λ is deviatoric (i.e., tr $\Lambda = 0$), the right-hand-side of (26), \mathcal{R} , 418 must comply with the restriction $tr(C\mathcal{R}) = 0$. This requires 419 \mathbb{T} to fulfil the condition tr[$C(\mathbb{T} : Y)$] = $C : \mathbb{T} : Y = 0$, for all 420 **Y**. 421

3.2 Remodelling laws

Equation (26) is the remodelling equation and it describes 423 how the anelastic phenomena evolve during all the defor-424 mative process. It is formulated as an evolution law for Π 425 through the tensor $\Lambda = \dot{\Pi} \Pi^{-1}$. 426

In this work, we assume that remodelling occurs at a 427 given material point when the Frobenius norm $\|\text{dev}\sigma\|$ = 428 $\sqrt{g_{ab}[\text{dev}\sigma]^{ac}g_{cd}[\text{dev}\sigma]^{db}}$ of the deviatoric part of the con-429 stitutive solid phase Cauchy stress, $\sigma = J^{-1} P F^{T}$, exceeds 430 at that point a threshold equivalent stress, σ_Y , termed "yield 431 stress" in analogy with Plasticity. To take this requirement 432 into account, we write \mathbb{T} as $\mathbb{T} = \zeta \mathbb{L}$, where ζ is a scalar stress-433 dependent "remodelling switch". Hence, following [31], we 434 prescribe a Perzyna-like model [51] 435

$$\zeta = \lambda(\phi_{\rm s}) \left[\frac{\|\text{dev}\boldsymbol{\sigma}\| - \sqrt{2/3}\,\sigma_Y}{\|\text{dev}\boldsymbol{\sigma}\|} \right]_+,\tag{27}$$

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where $\lambda(\phi_s)$ is a material parameter depending on the vol-436 umetric fraction of the solid phase, and the operator $[\cdot]_+$ 437 extracts the positive part of the function to which it is ap-438 plied (see also [8]). In this work, we assume that the yield 439 stress is constant, and we set $\sigma_Y = 0.002$ MPa. We emphasise 440 that $\lambda(\phi_s)$ vanishes for vanishing ϕ_s , since no remodelling 441 may occur if the solid phase is absent. In the following, we 442 adopt the simple law $\lambda(\phi_s) = \lambda_0 \phi_s^2 = \lambda_0 [\Phi_{s\nu}/JJ_{\Pi}]^2$, with 443 $\lambda_0 = 0.5 \ (\text{MPa} \cdot \text{s})^{-1}$. We also remark that, since the condi-444 tion $J_{\Pi} = 1$ applies in this context, the equality $\phi_s = \Phi_{s\nu}/J$ 445 allows to rephrase the dependence of λ on ϕ_s in terms of the 446 volume ratio J alone, rather than in terms of J and J_{Π} . 447

To complete the description of remodelling, it is necessary to specify the fourth-order tensor L. In this work, we consider the expression

$$\mathbb{L} = \mathbb{M}^* : \mathbb{D} : \mathbb{M}^{*\mathrm{T}},\tag{28}$$

where \mathbb{M}^* and \mathbb{M}^{*T} are specified in Appendix A. The fourthorder tensor \mathbb{D} encodes information about the material properties of the tissue and, in general, is a function of *C* and **II**. With the notation introduced in Appendix A, \mathbb{D} transforms tensors of $([T\mathcal{B}]_2^0, \text{sym})$ into tensors of $([T\mathcal{B}]_0^2, \text{sym})$. According to (28), the tensor \mathcal{R} featuring in (26) reads

$$\mathcal{R} = \mathbb{T} : Y = \zeta \mathbb{L} : Y = \zeta \mathbb{M}^* : \mathbb{D} : \mathbb{M}^{*\mathrm{T}} : Y.$$
⁽²⁹⁾

We remark that the double-contraction of \mathbb{M}^{*T} with Y extracts the deviatoric part of Y with respect to the metric C, i.e.,

$$\mathbb{M}^{*\mathrm{T}}: \mathbf{Y} = \mathbf{Y} - \frac{1}{3}\mathrm{tr}(\mathbf{C}^{-1}\mathbf{Y})\mathbf{C}.$$
(30)

⁴⁶⁰ Moreover, by introducing the tensor $Z := \mathbb{D} : \mathbb{M}^{*T} : Y$, the ⁴⁶¹ left-multiplication by \mathbb{M}^* in (29) leads to

$$\mathcal{R} = \zeta \,\mathbb{M}^* : \mathbf{Z} = \zeta \,\left[\mathbf{Z} - \frac{1}{3} \mathrm{tr}(\mathbf{C}\mathbf{Z})\mathbf{C}^{-1}\right],\tag{31}$$

⁴⁶² which guarantees the compliance with the constraint

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$$D = \operatorname{tr} \mathbf{\Lambda} = \operatorname{tr} \left[C \operatorname{sym}(\mathbf{\Lambda} C^{-1}) \right]$$
$$= -\operatorname{tr}(C\mathcal{R}) = -\zeta \operatorname{tr}[C(\mathbb{M}^* : \mathbf{Z})] = 0.$$
(32)

For the sake of simplicity, in the following we set $\mathbb{D} = \mathbb{I}^{\ddagger \ast}$ (see Appendix A for the definition of $\mathbb{I}^{\ddagger \ast}$), which implies

$$Z = \mathbb{I}^{\sharp *} : \mathbb{M}^{*T} : Y$$
$$= S - \frac{1}{2} \operatorname{tr}(CS) C^{-1} = \mathbb{M}^{*} : S = \tilde{S}, \qquad (33a)$$

$$\mathbb{L}: \boldsymbol{Y} = \mathbb{M}^*: \boldsymbol{Z} = \mathbb{M}^*: \mathbb{I}^{\sharp *}: \mathbb{M}^{*\mathrm{T}}: \boldsymbol{Y}$$
$$= \mathbb{M}^{\sharp *}: \boldsymbol{Y}, \tag{33b}$$

where \tilde{S} is said to be the deviatoric part of S with respect to the metric C, and $\mathbb{M}^{\sharp*}$ is defined in Appendix A. Furthermore, since \mathbb{M}^* is idempotent (i.e., it holds that $\mathbb{M}^* : \mathbb{M}^* = \mathbb{M}^*$), we obtain the identity

$$\mathbb{M}^* : \mathbf{Z} = \mathbb{M}^* : \mathbb{M}^* : \mathbf{S} = \mathbb{M}^* : \mathbf{S} = \mathbf{Z}.$$
 (34)

$$\mathcal{R} = \zeta \,\mathbb{M}^* : \mathbf{Z} = \zeta \,\mathbf{Z} = \zeta \left[\mathbf{S} - \frac{1}{3} \mathrm{tr}(\mathbf{CS}) \mathbf{C}^{-1} \right], \tag{35}$$

and the remodelling law takes on the form

$$\operatorname{sym}(\boldsymbol{\Lambda}\boldsymbol{C}^{-1}) = -\zeta \left[\boldsymbol{S} - \frac{1}{3}\operatorname{tr}(\boldsymbol{C}\boldsymbol{S})\boldsymbol{C}^{-1}\right] = -\zeta \,\tilde{\boldsymbol{S}},\tag{36}$$

thereby satisfying the requirement (32).

3.2.1 Model M1: Fully isotropic model 472

We use this model for comparison with the other ones, and we obtain it in the limit of vanishing volumetric fraction of the fibres. Hence, we set $\Phi_{1s\nu} = 0$, which implies $\Phi_{0s\nu} = \Phi_{s\nu}$, and we rewrite the strain energy density (10) as

$$\hat{W}_{\nu}(C_{\rm e}) = \Phi_{\rm s\nu}\hat{U}(J_{\rm e}) + \Phi_{\rm s\nu}\hat{W}_{0}(C_{\rm e}).$$
(37)

Consequently, the second Piola-Kirchhoff stress tensor consists of the isotropic contribution only, i.e., 477

$$S_{\rm iso} = \frac{1}{J_{\Pi}} \Pi \left[2\Phi_{\rm sv} \left(\frac{\partial \hat{U}}{\partial C_{\rm e}} + \frac{\partial \hat{W}_0}{\partial C_{\rm e}} \right) \right] \Pi^{\rm T}, \tag{38}$$

and the permeability tensor reduces to $K_{iso} = Jk_0C^{-1}$. Furthermore, we prescribe the remodelling law 480

$$\operatorname{sym}(\boldsymbol{\Lambda}\boldsymbol{C}^{-1}) = -\boldsymbol{\mathcal{R}}_{(1)} = -\boldsymbol{\zeta} \, \mathbb{L} : \boldsymbol{Y}_{\operatorname{iso}}. \tag{39}$$

with $Y_{iso} = CS_{iso}C$. By substituting Y_{iso} into (39) and performing all the necessary algebraic calculations, we obtain 482

$$\operatorname{sym}(\mathbf{\Lambda} \boldsymbol{C}^{-1}) = -\boldsymbol{\mathcal{R}}_{(1)} = -\boldsymbol{\zeta} \tilde{\boldsymbol{S}}_{\operatorname{iso}},\tag{40}$$

with
$$\tilde{\mathbf{S}}_{iso} = \mathbb{M}^*$$
: $\mathbf{S}_{iso} = \mathbf{S}_{iso} - \frac{1}{3} \operatorname{tr}(\mathbf{C}\mathbf{S}_{iso})\mathbf{C}^{-1}$.

3.2.2 Model M2: Semi-isotropic model

In this model, we use the full permeability tensor defined in (20) and the transversely isotropic strain energy density (10), which produces the second Piola-Kirchhoff stress tensor

$$\mathbf{S} = \mathbf{S}_{\mathrm{i}} + \mathbf{S}_{\mathrm{a}},\tag{41}$$

with

$$\mathbf{S}_{i} = \frac{1}{J_{\Pi}} \mathbf{\Pi} \left[2\Phi_{sv} \frac{\partial \hat{U}}{\partial C_{e}} + 2\Phi_{0sv} \frac{\partial \hat{W}_{0}}{\partial C_{e}} + 2\Phi_{1sv} \frac{\partial \hat{W}_{1i}}{\partial C_{e}} \right] \mathbf{\Pi}^{\mathrm{T}},$$
(42a)

$$\boldsymbol{S}_{a} = \frac{1}{J_{\Pi}} \boldsymbol{\Pi} \left[2 \Phi_{1s\nu} \frac{\partial \langle \hat{W}_{1a} \rangle}{\partial \boldsymbol{C}_{e}} \right] \boldsymbol{\Pi}^{\mathrm{T}}.$$
 (42b)

Note that S_i and S_a represent, respectively, the isotropic and transversely isotropic contributions to the overall constitutive part of the second Piola-Kirchhoff stress tensor of the solid phase, S.

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In spite of the fact that both the elastic and the hydraulic
 response of the tissue are transversely isotropic, we consider
 the same remodelling law as in the Model M1. Hence, we set

$$\operatorname{sym}(\mathbf{\Lambda} \mathbf{C}^{-1}) = -\mathcal{R}_{(2)} = -\zeta \,\mathbb{L} : \mathbf{Y},\tag{43}$$

where *Y* splits additively as $Y = CSC = CS_iC + CS_aC$. Analogously to the Model M1, also in this case the remodelling law can be written as

$$\operatorname{sym}(\boldsymbol{\Lambda}\boldsymbol{C}^{-1}) = -\boldsymbol{\mathcal{R}}_{(2)} = -\boldsymbol{\zeta}\left[\boldsymbol{\tilde{S}}_{i} + \boldsymbol{\tilde{S}}_{a}\right],\tag{44}$$

500 with

$$\tilde{\boldsymbol{S}}_{i} = \boldsymbol{\mathbb{M}}^{*} : \boldsymbol{S}_{i} = \boldsymbol{S}_{i} - \frac{1}{3} \operatorname{tr}(\boldsymbol{C}\boldsymbol{S}_{i})\boldsymbol{C}^{-1},$$
(45a)

$$\tilde{\boldsymbol{S}}_{a} = \mathbb{M}^{*} : \boldsymbol{S}_{a} = \boldsymbol{S}_{a} - \frac{1}{3} \operatorname{tr}(\boldsymbol{C}\boldsymbol{S}_{a})\boldsymbol{C}^{-1}$$
(45b)

⁵⁰¹ being the deviatoric parts of S_i and S_a , respectively, with ⁵⁰² respect to the deformed metric *C*. We remark that, according ⁵⁰³ to (44), the presence of the fibres supplies a direct contribu-⁵⁰⁴ tion to the remodelling law through \tilde{S}_a .

Remark Each remodelling law, i.e., (39) or (43), is in general 505 equivalent to a set of six scalar differential equations in the 506 components of Π . However, when the isochoric condition 507 $J_{\Pi} = 1$ is enforced, as is the case in this work, the num-508 ber of independent equations is five, because the constraint 509 $tr(\Lambda) = tr(\dot{\Pi}\Pi^{-1}) = 0$ has to be respected. Since, in general, 510 Π possesses nine independent components, which become 511 eight when the isochoric condition $J_{\Pi} = 1$ applies, the re-512 modelling laws are not closed. To obtain the closure, we per-513 form the polar decomposition of Π , i.e., $\Pi = V.R \equiv VGR$, 514 where R is a rotation tensor and V is a symmetric and 515 positive-definite tensor. In this work, we impose that the ro-516 tations associated with remodelling are not allowed, so that 517 only V is unknown. Since it has only six independent com-518 ponents (actually five, because it holds that $J_{\Pi} = \det V = 1$), 519 the remodelling laws become closed. We also notice that the 520 identity $\Lambda = \dot{\Pi} \Pi^{-1} = \dot{V} V^{-1}$ holds true. 521

522 4 Benchmark test and numerical settings

We formulate a finite strain poroplastic problem for a porous medium in which the interstitial fluid obeys Darcy's law and the solid phase exhibits hyperelastic behaviour. Given the reference configuration of the tissue $\mathcal{B} \subset S$ and the interval of time $\mathcal{I} \subset \mathbb{R}$, find the motion χ , pressure p, and V such that

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	\mathbf{n} Urau $\boldsymbol{\nu}$	J = J.	$\Pi D \wedge I$.	(4 0a)
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$$\operatorname{Div}\left(-Jp\,\boldsymbol{g}^{-1}\boldsymbol{F}^{-\mathrm{T}}+\boldsymbol{P}\right)=\boldsymbol{0},\qquad \text{in }\mathcal{B}\times\boldsymbol{I},\qquad (46b)$$

$$\operatorname{sym}(\mathbf{\Lambda} \mathbf{C}^{-1}) = -\mathbf{\mathcal{R}},$$
 in $\mathcal{B} \times \mathbf{I},$ (46c)

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where \mathcal{R} can be equal to $\mathcal{R}_{(1)}$ or $\mathcal{R}_{(2)}$, depending on whether 529 the model M1 or M2 is computed. We emphasise that, by con-530 struction, both $\mathcal{R}_{(1)}$ and $\mathcal{R}_{(2)}$ have to be understood as func-531 tionals of χ , and V, i.e., $\mathcal{R}_{(\alpha)} = \hat{\mathcal{R}}_{(\alpha)}(\chi, V)$, for $\alpha \in \{1, 2\}$. 532 Whereas (46c) expresses the general form of the investigated 533 remodelling law, (46a) and (46b) represent, respectively, the 534 mass balance law and the momentum balance law for the 535 biphasic system with which the tissue is approximated. We 536 recall, indeed, that the tissue is assumed here to consist of 537 a solid phase, which comprises a porous matrix and the 538 reinforcing fibres, and an inviscid interstitial fluid obeying 539 Darcy's law. Equations (46a)-(46c) are determined under 540 the hypotheses that the mass densities of the solid and the 541 fluid phase are constant (a condition implying the intrinsic 542 incompressibility of both phases), and that all the external 543 544 the quantities of order higher than the first in the relative 545 velocity $v_{\rm fs} := v_{\rm f} - v_{\rm s}$ are negligible. More specifically, the 546 mass balance law (46a) implies that the opposite of the di-547 vergence of the specific (material) mass flux Q = -KGrad p 548 is compensated for by the time derivative of the volume ratio 549 J. Furthermore, the momentum balance law (46b) defines 550 the overall stress tensor of the biphasic system under study 551 as $P_{\text{tot}} = -Jp \, g^{-1} F^{-T} + P$, where the pressure p is the La-552 grange multiplier associated with the incompressibility and 553 the saturation constraints. 554

The logical steps leading to (46a) and (46b) have been presented elsewhere (cf. e.g. [37, 19, 61, 33, 38, 39, 5, 40]), and will not be repeated here. In addition to them, the remodelling law (46c) supplies a further coupling among deformation, pressure, and plastic-like distortions.

Equations (46a)–(46c) shall be solved for simulating an 560 unconfined compression test of the sample under study. This 561 test represents a typical benchmark problem for investigating 562 the elastic and hydraulic properties of biological tissues (cf. 563 (46a) and (46b), respectively), and has been adapted here in 564 order to also account for the reorganisation of the sample's 565 internal structure (cf. (46c)). In the experiment simulated in 566 this work, a specimen of tissue of cylindric shape is posi-567 tioned between two rigid, parallel plates, and compressed. 568 The two plates are impermeable to the fluid flow. The com-569 pression takes place in displacement control and, in particu-570 lar, by displacing the upper plate according to a given loading 571 ramp. The lower plate is instead kept fixed, and the specimen 572 is clamped on it. The upper plate constitutes a frictionless 573 glide surface for the specimen, whose upper boundary is thus 574 allowed to deform radially in axial-symmetric way. The lat-575 eral boundary is assumed to be free of contact forces, thereby 576 requiring that both the pressure and the radial component of 577 the overall stress vanish on it (see (47b)). 578

By introducing a reference frame with origin *O* coinciding with the centre of the lower boundary of the sample, and orthonormal cartesian basis vectors $\{\Xi_I\}_{I=1}^3$ emanating from *O*, such that Ξ_3 is the unit vector directed along the specimen's symmetry axis, the experiment described above is represented by the boundary conditions [5,34]:

$$\begin{cases} \chi^3 = f \\ (-K \operatorname{Grad} p) \cdot N = 0 \end{cases} \quad \text{on } \partial \mathcal{B}^{(u)}, \qquad (47a)$$

$$\begin{cases} (-Jpg^{-1}F^{-T} + P).N = 0\\ p = 0 \end{cases} \quad \text{on } \partial \mathcal{B}^{(l)}, \tag{47b}$$

$$\begin{cases} \chi(X,t) - \chi(X,0) = \mathbf{0} \\ (-K \operatorname{Grad} p).N = 0 \end{cases} \quad \text{on } \partial \mathcal{B}^{(L)}. \tag{47c}$$

In (47a), χ^3 is the axial component of the motion, and f is the loading ramp

$$f(t) = \begin{cases} L - \frac{t}{T_{\text{ramp}}} u_{\text{T}}, & \text{for } t \in [0, T_{\text{ramp}}], \\ L - u_{\text{T}}, & \text{for } t \in [T_{\text{ramp}}, T_{\text{end}}], \end{cases}$$
(48)

where $u_{\rm T} = 0.20$ mm is the target displacement imposed to 587 the sample and L = 1 mm is the sample's initial length. The 588 initial cross section of the sample has diameter D = 3 mm. 589 The target displacement is reached at the end of the loading 590 ramp, i.e., at $T_{\text{ramp}} = 20$ s, and is then kept constant until 591 $T_{\text{end}} = 300 \text{ s. Moreover, in (47a)-(47c), } \partial \mathcal{B}^{(u)}, \ \partial \mathcal{B}^{(l)} \text{ and}$ 592 $\partial \mathcal{B}^{(L)}$ are the upper, lateral and lower part of the boundary 593 $\partial \mathcal{B}$, such that $\partial \mathcal{B} = \partial \mathcal{B}^{(u)} \sqcup \partial \mathcal{B}^{(l)} \sqcup \partial \mathcal{B}^{(L)}$. Finally, N is the 594 unit vector normal to $\partial \mathcal{B}$. 595

It is assumed that, at the initial time, the sample finds itself in an undeformed state, with zero pressure, and in the absence of anelastic distortions. These requirements lead to the initial conditions

$$\chi(X,0) = X, \qquad \forall X \in \mathcal{B}, \qquad (49a)$$

$$p(X,0) = 0, \qquad \forall X \in \mathcal{B}, \qquad (49b)$$

$$V(X,0) = G^{-1}(X) \qquad \forall X \in \mathcal{B}.$$
(49c)

The numerical solution of (46a)-(46c), with (49a)-(49c) 600 and (47a)-(47c), is achieved by performing Finite Element 601 simulations. In particular, following [39,5], (46a) and (46b) 602 are put in weak form, and solved according to a given Finite 603 Element scheme, while (46c) is solved only at the integra-604 tion points of the finite elements. To this end, by search-605 ing for the motion χ and pressure p in the Sobolev spaces 606 $(H^1(\mathcal{B} \times \mathcal{I}, \mathcal{S}))^3$ and $H^1(\mathcal{B} \times \mathcal{I}, \mathcal{S})$, respectively, and enforc-607 ing the boundary conditions (47a)-(47c), the model equa-608 tions (46a)-(46c) are reformulated as 609

$$\mathcal{F}_{\chi} = \hat{\mathcal{F}}_{\chi}(\chi, p, V)$$
$$= \int_{\mathcal{B}} \hat{P}(\chi, p, V) : \boldsymbol{g} \operatorname{Grad} \tilde{\boldsymbol{u}} = 0,$$
(50a)

$$\mathcal{F}_{p} = \hat{\mathcal{F}}_{p}(\chi, p, V)$$
$$= \int_{\mathcal{B}} \left\{ (\operatorname{Grad} \tilde{p}) \hat{K}(\chi, V) (\operatorname{Grad} p) + \tilde{p} \dot{J} \right\} = 0, \quad (50b)$$

$$\mathcal{F}_{V} = \hat{\mathcal{F}}_{V}(\chi, V) = \operatorname{sym}(\dot{V}V^{-1}) + \hat{\mathcal{R}}(\chi, V) = \mathbf{0},$$
 (50c)

where $\tilde{\boldsymbol{u}}$ and \tilde{p} are the test functions associated with the velocity and pressure and are sometimes referred to as "virtual velocity" and "virtual pressure", respectively. We notice that the functionals $\hat{\mathcal{F}}_{\chi}$ and $\hat{\mathcal{F}}_{p}$ depend linearly on the virtual fields $\tilde{\boldsymbol{u}}$ and \tilde{p} . However, for the sake of a lighter notation, we have omitted this dependence in their definitions.

5 Results

In this section, we present and discuss the main results of 617 our simulations (see Figures 2–6). In particular, we show (i) 618 how remodelling modulates the mechanical and hydraulic 619 response of the tissue, and (ii) how the fibre reinforcement, 620 which makes the tissue transversely isotropic, influences the 621 evolution of the anelastic distortions. To highlight the conse-622 quences of remodelling, we run a set of simulations in which 623 remodelling is switched off, and we compare the correspond-624 ing results with those stemming from the set of simulations 625 in which the models M1 and M2 are implemented. Moreover, 626 in order to see the role played by the fibre reinforcement, we 627 compare the results predicted by the model M1, in which an 628 ideal isotropic tissue without fibres is simulated, with those 629 predicted by the model M2, in which the presence of the 630 fibres is accounted for. In all the plots (Figures 2-6), we 631 evaluate the physical quantity of interest at the point $X_{\rm U}$ of 632 Cartesian coordinates given by (1.3, 0.0, 1.0) [mm], which is 633 on the upper boundary and close to the lateral boundary of 634 the sample. 635

In Figure 2a, we report the time trend of the magni-636 tude of the (spatial) filtration velocity, $\|\boldsymbol{q}(X_{\rm U}, t)\|$, evaluated 637 at the point X_U for $t \in [0, T_{end}]$, where we let T_{end} be ar-638 bitrarily greater than T_{ramp} . We show both the case of no 639 remodelling and the case of remodelling, as described by the 640 models M1 and M2. In the absence of remodelling, the mag-641 nitude of the filtration velocity grows monotonically until 642 the target displacement is reached, i.e., until $t = T_{ramp}$. Then, 643 it relaxes asymptotically towards zero for increasing time. 644 When remodelling occurs, the trend of $||q(X_U, t)||$ depends 645 on whether or not the fibres are accounted for. In the sim-646 ulation performed by applying the model M1, the influence 647 of remodelling on $\|\boldsymbol{q}(X_{\mathrm{U}},t)\|_{\mathrm{M1}}$ is twofold: on the one hand, 648 it lowers considerably the maximum value of $\|\boldsymbol{q}(X_{\mathrm{U}},t)\|$, 649 which is however attained at $t = T_{ramp}$, and, on the other 650 hand, it leads to a much slower relaxation time. Hence, even 651 though $\|\boldsymbol{q}(X_{\mathrm{U}},t)\|_{\mathrm{M1}}$ decreases monotonically towards zero, 652 the curve associated with M1 intersects the curve of no re-653 modelling, and it holds that 654

$$\|\boldsymbol{q}(X_{\mathrm{U}},t)\|_{\mathrm{M1}} \ge \|\boldsymbol{q}(X_{\mathrm{U}},t)\|_{\mathrm{no-rem}},$$
(51)

for all $t \ge T_1$, with $T_1 > T_{ramp}$ being the time at which the two curves intersect each other. The simulation performed considering the model M2 leads, instead, to quite different

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50

100

0.005

0.000



Fig. 2 Norm of Darcy velocity vs time (a) and radial component of Darcy velocity vs time (b), evaluated at the point X_U of Cartesian coordinates (1.3, 0.0, 1.0) [mm]. The anisotropic model predicts an inversion of the filtration velocity, which yields thus an inflow of fluid after a critical instant of time is reached. This behaviour is not captured by the isotropic model M1.

results. First of all, the maximum value of $\|\boldsymbol{q}(X_{\mathrm{U}},t)\|_{\mathrm{M2}}$, al-658 ways attained at $T = T_{ramp}$, is smaller than the one reached in 659 the case of no remodelling and bigger than the one predicted 660 by M1. Moreover, the relaxation of $\|\boldsymbol{q}(X_{\mathrm{U}},t)\|_{\mathrm{M2}}$ towards 661 zero is slower than that observed in the case of no remod-662 elling, but slightly faster than the one obtained by employing 663 the model M1. The most noticeable results, however, are 664 given by the loss of monotonicity of $\|\boldsymbol{q}(X_{\mathrm{U}},t)\|_{\mathrm{M2}}$ in the 665 interval $[T_{ramp}, T_{end}]$, and by the presence of the point of 666 non-differentiability, herafter denoted by T_c , between T_{ramp} 667 and t = 50 s. This behaviour is due to the fact that, when 668 remodelling occurs and the anisotropy of the fibre pattern 669 is considered, the radial component of the filtration velocity 670 decreases for $t > T_{ramp}$, becomes negative until it attains a 671 global minimum and, subsequently, it grows asymptotically 672 towards zero for a sufficiently long time (see Figure 2b). 673

The change of sign in the radial velocity may be interpreted as a "syringe effect", thereby meaning that, for $t > T_c$, the fluid tends to flow back into the tissue. Since the fluid filtration velocity complies with Darcy's law, this behaviour is accompanied by a change of sign of the radial pressure gradient, which implies that the pressure at X_U becomes



150 Time [s] 200

smaller than zero for $t > T_c$ (we recall, indeed, that our 680 boundary conditions prescribe that, on the lateral bound-681 ary of the sample, the pressure is zero at all times). This 682 observation seems to be supported by the results shown in 683 Figure 3. In the absence of remodelling, pressure grows until 684 a global maximum is reached, and it relaxes then towards 685 zero for increasing time. A qualitatively similar trend is also 686 observed when remodelling is switched on and the model M1 687 is used, even though the maximum value of pressure is much 688 smaller than the one obtained in the case of no remodelling. 689 The model M1 predicts, indeed, that $[p(X_U, t)]_{M1}$ consists 690 of two monotonic branches, one increasing over the inter-691 val $[0, T_{ramp}]$ and the other one decreasing over $[T_{ramp}, T_{end}]$. 692 The decreasing branch intersects the relaxing branch of the 693 pressure curve of no remodelling and tends towards zero 694 more slowly than the latter one. The curve determined by 695 simulating the model M2 grows rather steeply until the max-696 imum pressure is attained, and this maximum places itself 697 in between the values obtained in the case of no remdelling 698 and that of the model M1, respectively. Then, $[p(X_U, t)]_{M2}$ 699 decreases much faster than it happens in the other cases, be-700 comes negative, and reaches a global minimum. Afterwards 701 it grows again, and it then tends to zero from below at a 702 rate comparable with that of no remodelling. We remark that 703 the instant of time at which pressure equals zero coincides 704 with T_c , i.e., the time at which the radial component of the 705 filtration velocity changes its sign. 706

In Figure 4, we study the time trend of porosity at $X_{\rm U}$. 707 We notice that, both in the case of no remodelling and in 708 the case of the model M1, porosity decreases monotonically 709 in time. In the absence of remodelling, porosity varies very 710 smoothly, and the amplitude of the variation between its ini-711

No Remodel

Model M1 Model M2

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Fig. 4 Porosity vs time, $1 - \Phi_{sv}(X_U)/J(X_U, t)$, evaluated at the point X_U of Cartesian coordinates (1.3, 0.0, 1.0) [mm]. Whereas the case of no remodelling and the model M1 predict quantitatively different, but qualitatively similar, results, the model M2 is characterised by a trend that is both quantitatively and qualitatively different from the other two. The loss of monotonicity is, in fact, consistent with that of Figures 2 and 3, and represents an opening of the tissue's pores (with corresponding increase of porosity) on the way towards the stationary state.

tial and asymptotic values is bigger than in the other case. 712 The model M1, in turn, predicts a rather pronounced change 713 of slope of the porosity curve, and the asymptotic value of 714 porosity is reached more slowly. A quite different behaviour 715 can be observed when the tissue's anisotropy is accounted 716 for. Indeed, in accordance with the inversion of the fluid fil-717 tration velocity (see Figure 2) and the change of sign of the 718 pressure (see Figure 3), the model M2 prescribes that poros-719 ity varies in time in a non-monotonic way. More specifically, 720 it decreases until it comes to a global minimum, which cor-721 responds to the end of the loading ramp, and then it grows 722 towards a stationary value. This behaviour is consistent with 723 the fact that, to permit the inflow of fluid, the tissue must 724 increase its porosity, and it seems to be a consequence of the 725 interplay between the tissue's anisotropy and the evolution 726 of the anelastic distortions. 727

⁷²⁸ In terms of $F_{\rm p}$, a measure of the magnitude of plastic-⁷²⁹ like distortions is the Frobenius norm of the anelastic strain ⁷³⁰ tensor

$$\boldsymbol{E}_{\mathrm{p}} = \frac{1}{2} \left[\boldsymbol{F}_{\mathrm{p}}^{\mathrm{T}} \cdot \boldsymbol{F}_{\mathrm{p}} - \boldsymbol{G} \right].$$
(52)

⁷³¹ Since it holds that F_p is the inverse of Π , E_p may be rewritten ⁷³² as

$$\boldsymbol{E}_{\mathrm{p}} = \frac{1}{2} [\boldsymbol{\Pi}^{-\mathrm{T}} \cdot \boldsymbol{\Pi}^{-1} - \boldsymbol{G}] = -\boldsymbol{\mathcal{A}}_{\boldsymbol{\Pi}},$$
(53)

⁷³³ where \mathcal{A}_{Π} is the Almansi-Euler-like strain tensor associated ⁷³⁴ with Π . Finally, by enforcing the polar decomposition $\Pi =$ ⁷³⁵ *V*.*R*, *E*_p becomes

$$E_{\rm p} = \frac{1}{2} [V^{-1} \cdot V^{-1} - G].$$
(54)



Fig. 5 Frobenius norm of $E_p = \frac{1}{2}[V^{-1}.V^{-1}-G]$ vs time, $||E_p(X_U, t)||$, evaluated at the point X_U of Cartesian coordinates (1.3, 0.0, 1.0) [mm]. The magnitude of the plastic strains is bigger in the transversely isotropic model M2. In the isotropic model M1, instead, the plastic strains are rather small, but they tend to the stationary state much more slowly than predicted by the model M2.

Equation (53) suggests which tensor field should be used to address remodelling within the theory of uniformity [15,50, 7,54].

The Frobenius norm of E_p is now evaluated at X_U and 739 its variation in time is reported in Figure 5. We notice that 740 the magnitude of the anelastic distortions as predicted by the 741 model M2 is much bigger than that obtained by the model 742 M1. Thus, the anisotropy of the tissue seems to enhance the 743 growth of the plastic distortions, whose magnitude increases 744 quite rapidly and tends to approach a stationary value. In 745 the case of the model M1, instead, $||E_p(X_U, t)||$ grows much 746 more slowly (and almost linearly) towards a stationary value. 747



Fig. 6 Equivalent stress vs time, evaluated at the point X_U of Cartesian coordinates (1.3, 0.0, 1.0) [mm]. The equivalent stress is the Frobenius norm of the deviatoric part of the constitutive Cauchy stress tensor, i.e., $\|\det \sigma\|$, with $\sigma = J^{-1}FSF^{T}$. The 2nd Piola-Kirchhoff stress tensor *S* is given by (38) for the model M1, and by (41) both for the model M2 and for the case of no remodelling (in which, however, the identity $V = G^{-1}$ applies).

Finally, we investigate how the onset of plastic distortions 748 modulates the stress borne by the tissue. To this end, we plot 749 in Figure 6 the von Mises equivalent stress at $X_{\rm U}$, and we 750 notice that the curve corresponding to the model M1 is, until 751 about 200 s, bounded from above by the curve pertaining 752 to the model M2. This means that, even though the plastic 753 distortions are characterised by a magnitude of $E_{\rm p}$ that is 754 bigger in the anisotropic case than in the isotropic one, the 755 level of stress reached in the first case is higher. We remark 756 that the onset of remodelling occurs only when the von Mises 757 equivalent stress, $\|\text{dev}\sigma\|$, overcomes the yield stress, σ_Y . In 758 fact, there exists an instant of time such that the condition 759 of incipient remodelling, i.e., $\|\text{dev}\boldsymbol{\sigma}\| = \sigma_Y$, is verified, and 760 the von Mises equivalent stress is bigger than σ_Y for all 761 subsequent times. To highlight this behaviour, we plotted in 762 Figure 6 the yield stress (which is constant in time in this 763 work), and we showed that, in all the considered cases, the 764 von Mises equivalent stress exceeds the yield stress after a 765 quite short interval of time. 766

67 6 Conclusions

In this work, we employed an inhomogeneous and transversely isotropic poroplastic model of fibre-reinforced biological tissue in order to study how the variation of the tissue's internal structure (i.e., the process of remodelling), which manifests itself through the onset and evolution of anelastic distortions, is influenced by the material symmetries of the tissue itself.

For our purposes, we rephrased the poroelastic model 775 of hydrated, fibre-reinforced tissues summarised in [19,61] 776 in order to account for the presence of anelastic distortions 777 (the definition of the hyperelastic strain energy energy is 778 developed from [24, 19] and the tissue's permeability has 779 been adapted from [23, 22, 19, 61]). Then, we formulated and 780 solved numerically the two different descriptions of struc-781 tural remodelling denoted by model M1 and model M2. We 782 recall that, while the tissue has been simulated as inhomo-783 geneous and transversely isotropic both in the case of the 784 model M2 and in the reference case of no remodelling, it 785 has been regarded as inhomogeneous but isotropic in the 786 model M1. We emphasise that this idealisation serves as a 787 basis for comparison with the transversely isotropic model 788 M2, and has been done to highlight the interplay between the 789 tissue's material symmetries and the development of plastic 790 distortions. These, indeed, drive an evolution of the group of 791 material symmetries, but they do not change it (see [12,13] 792 for further details). 793

Among the obtained results, represented graphically in Figures 2–6, we give prominence to the "syringe effect" discussed in Section 5, which is observed in our simulations only when remodelling occurs in the tissue modelled as an inhomogeneous and tranversely isotropic material (cf. model M2). Such effect seems to be an evidence of the change of the tissue's mechanical and hydraulic behaviour. Such alteration of material response could characterise a diseased or damaged tissue, and could thus also provide some indications on how the tissue might behave in non-physiological conditions.

Finally, since the observed changes of material behaviour 805 occurs both qualitatively and quantitatively in the case of 806 anisotropy (while the change is only quantitative in the case 807 of isotropy), our results could be used for studying the inter-808 play between growth and remodelling in anisotropic tissues. 809 For example, this could be of interest for elaborating more de-810 tailed and more accurate models of tumour growth, in which 811 the onset of remodelling has appreciable consequences on 812 the tumour evolution [49,48]. 813

Appendix A: Fourth-order tensors

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The notation adopted in the following is taken from [17]. Let $[T\mathcal{B}]_1^1, [T\mathcal{B}]_1^1, [T\mathcal{B}]_0^2$, and $[T\mathcal{B}]_2^0$ denote the spaces of all second-order tensors which, as bilinear maps, read ⁸¹⁷

- $A: T^{\star}\mathcal{B} \times T\mathcal{B} \to \mathbb{R}, \tag{55a}$
- $\boldsymbol{B}: T\mathcal{B} \times T^{\star}\mathcal{B} \to \mathbb{R}, \tag{55b}$

$$\boldsymbol{T}: T^{\star}\mathcal{B} \times T^{\star}\mathcal{B} \to \mathbb{R}, \tag{55c}$$

$$Q: T\mathcal{B} \times T\mathcal{B} \to \mathbb{R},\tag{55d}$$

respectively. Let also $([T\mathcal{B}]_0^2, \text{sym})$ and $([T\mathcal{B}]_2^0, \text{sym})$ be, respectively, the subspaces of $[T\mathcal{B}]_0^2$ and $[T\mathcal{B}]_2^0$ of all symmetric, second-order tensors. The elements of $[T\mathcal{B}]_1^1$ and $[T\mathcal{B}]_1^1$ and $[T\mathcal{B}]_1^1$ can be written as linear maps from $T\mathcal{B}$ into itself, and from $T^*\mathcal{B}$ into itself, respectively, while the elements of $[T\mathcal{B}]_0^2$, and $[T\mathcal{B}]_2^0$ can be written as linear maps from $T^*\mathcal{B}$ into $T\mathcal{B}$, and from $T\mathcal{B}$ into $T^*\mathcal{B}$, respectively.

Let us also consider the spaces $[T\mathcal{B}]_2^2$ and $[T\mathcal{B}]_2^2$ of all fourth-order tensors of the type

$$\mathbb{T} \in [T\mathcal{B}]_{2}^{2}, \ \mathbb{T} : T^{\star}\mathcal{B} \times T^{\star}\mathcal{B} \times T\mathcal{B} \times T\mathcal{B} \to \mathbb{R}, \\ \mathbb{Q} \in [T\mathcal{B}]_{2}^{2}, \ \mathbb{Q} : T\mathcal{B} \times T\mathcal{B} \times T^{\star}\mathcal{B} \times T^{\star}\mathcal{B} \to \mathbb{R}.$$

An element of $[T\mathcal{B}]_2^2$ can also be represented as a linear map from $[T\mathcal{B}]_0^2$ into $[T\mathcal{B}]_0^2$. Analogously, an element of $[T\mathcal{B}]_2^2$ can be represented as a linear map from $[T\mathcal{B}]_2^0$ into $[T\mathcal{B}]_2^0$. For instance, the fourth-order tensor

$$\mathbb{I} : [T\mathcal{B}]_0^2 \to ([T\mathcal{B}]_0^2, \operatorname{sym}), \\
\mathbb{I} = \frac{1}{2} \left(\mathbf{I} \otimes \mathbf{I} + \mathbf{I} \overline{\otimes} \mathbf{I} \right),$$
(57)

where $\boldsymbol{I}: T\mathcal{B} \to T\mathcal{B}$ is the identity tensor in $T\mathcal{B}$, returns the symmetric part of the element of $[T\mathcal{B}]_0^2$ to which it is applied. Given two tensors $\boldsymbol{A}, \boldsymbol{D} \in [T\mathcal{B}]_1^1$, the representation of the tensor products $\boldsymbol{A} \otimes \boldsymbol{D}$ and $\boldsymbol{A} \otimes \boldsymbol{D}$ in index notation reads $[\boldsymbol{A} \otimes \boldsymbol{D}]_{MN}^{AB} = A_M^A \overline{D}_N^B$ and $[\boldsymbol{A} \otimes \boldsymbol{D}]_{MN}^{AB} = A_N^A D_M^B$ ⁸³⁵

[9]. Accordingly, in index notation, I is represented by the 836 expression 837

$$\mathbb{I}^{AB}_{\ MN} = \frac{1}{2} \left(\delta^A_{\ M} \delta^B_{\ N} + \delta^A_{\ N} \delta^B_{\ M} \right). \tag{58}$$

Thus, for every $T \in [T\mathcal{B}]_0^2$, it holds that 838

$$\mathbb{I}: \boldsymbol{T} = \frac{1}{2} \left(\boldsymbol{T} + \boldsymbol{T}^{\mathrm{T}} \right) = \operatorname{sym}(\boldsymbol{T}),$$
(59)

where the symbol ":" stands for "double contraction". In in-839 dex notation, it reads $(\mathbb{I}: T)^{AB} = \mathbb{I}^{AB}_{MN} T^{MN} = [\text{sym}(T)]^{AB}$. 840 By definition, I is the identity fourth-order tensor over the 841 space $([T\mathcal{B}]_{0}^{2}, \text{sym})$. From here on, we consider only the re-842 strictions of the fourth-order tensors of $[T\mathcal{B}]_0^2$ onto $([T\mathcal{B}]_0^2$, sym). 843 For every $T \in ([T\mathcal{B}]_0^2, \text{sym})$, the fourth-order tensor 844

$$\mathbb{K}^* : ([T\mathcal{B}]^2_0, \operatorname{sym}) \to ([T\mathcal{B}]^2_0, \operatorname{sym}),$$
$$\mathbb{K}^* = \frac{1}{3} C^{-1} \otimes C$$
(60)

extracts the spherical part of T with respect to the metric C, 845 i.e., 846

$$\mathbb{K}^*: T = \frac{1}{2} \operatorname{tr}(CT) C^{-1}.$$
(61)

The deviatoric part of T with respect to the metric C is 847 obtained by substracting \mathbb{K}^* : *T* to *T*. This operation can be 848 represented by the application of the fourth-order tensor 849

$$\mathbb{M}^* : ([T\mathcal{B}]_0^2, \operatorname{sym}) \to ([T\mathcal{B}]_0^2, \operatorname{sym})$$
$$\mathbb{M}^* = \mathbb{I} - \mathbb{K}^*, \tag{62}$$

to T i.e.,

$$\mathbb{M}^*: \boldsymbol{T} = (\mathbb{I} - \mathbb{K}^*): \boldsymbol{T} = \boldsymbol{T} - \frac{1}{3} \operatorname{tr}(\boldsymbol{CT}) \boldsymbol{C}^{-1}.$$
(63)

Clearly, it holds that tr $[C(\mathbb{M}^* : T)] = 0$. We remark that, by 85 their own definition, \mathbb{K}^* and \mathbb{M}^* constitute the partition of 852 unity, i.e., $\mathbb{I} = \mathbb{K}^* + \mathbb{M}^*$. 853

In analogous manner, we introduce the identity fourth-854 order tensor over the space $([T\mathcal{B}]_2^0, \text{sym})$, i.e., 855

$$\mathbb{I}^{\mathrm{T}} : ([T\mathcal{B}]_{2}^{0}, \operatorname{sym}) \to ([T\mathcal{B}]_{2}^{0}, \operatorname{sym}),$$
$$\mathbb{I}^{\mathrm{T}} = \frac{1}{2} \left(\boldsymbol{I}^{\mathrm{T}} \underline{\otimes} \boldsymbol{I}^{\mathrm{T}} + \boldsymbol{I}^{\mathrm{T}} \overline{\otimes} \boldsymbol{I}^{\mathrm{T}} \right), \tag{64}$$

where $I^{\mathrm{T}}: T^{\star}\mathcal{B} \to T^{\star}\mathcal{B}$ is the identity tensor in $T^{\star}\mathcal{B}$. For 856 every $\boldsymbol{Q} \in ([T\mathcal{B}]_2^0, \text{sym})$ it holds that 857

$$\mathbb{I}^{\mathrm{T}}: \boldsymbol{Q} = \frac{1}{2} \left(\boldsymbol{Q} + \boldsymbol{Q}^{\mathrm{T}} \right) \equiv \boldsymbol{Q}.$$
(65)

The spherical and the deviatoric parts of Q with respect 858 to the inverse metric C^{-1} are extracted by employing the 859 fourth-order tensors 860

$$\mathbb{K}^{*T} : ([T\mathcal{B}]_2^0, \operatorname{sym}) \to ([T\mathcal{B}]_2^0, \operatorname{sym}),$$
$$\mathbb{K}^{*T} = \frac{1}{3} \mathcal{C} \otimes \mathcal{C}^{-1}, \tag{66}$$

and

$$\mathbb{M}^{*\mathrm{T}} : ([T\mathcal{B}]_2^0, \mathrm{sym}) \to ([T\mathcal{B}]_2^0, \mathrm{sym}),$$
$$\mathbb{M}^{*\mathrm{T}} = \mathbb{I}^{\mathrm{T}} - \mathbb{K}^{*\mathrm{T}},$$

respectively, which are such that

$$\mathbb{K}^{*\mathrm{T}}: \boldsymbol{Q} = \frac{1}{3} \mathrm{tr}(\boldsymbol{C}^{-1}\boldsymbol{Q})\boldsymbol{C}, \tag{68}$$

$$\mathbb{M}^{*\mathrm{T}}: \boldsymbol{Q} = (\mathbb{I}^{\mathrm{T}} - \mathbb{K}^{*\mathrm{T}}): \boldsymbol{Q} = \boldsymbol{Q} - \frac{1}{3} \mathrm{tr}(\boldsymbol{C}^{-1}\boldsymbol{Q})\boldsymbol{C}.$$
(69)

In this case, it holds that tr $[C^{-1}(\mathbb{M}^{*T} : Q)] = 0.$ 863 Finally, we introduce the fourth-order tensor 864

$$\mathbb{I}^{\sharp*} : ([T\mathcal{B}]_2^0, \operatorname{sym}) \to ([T\mathcal{B}]_0^2, \operatorname{sym}),$$
$$\mathbb{I}^{\sharp*} = \frac{1}{2} \left(\boldsymbol{C}^{-1} \underline{\otimes} \, \boldsymbol{C}^{-1} + \boldsymbol{C}^{-1} \, \overline{\otimes} \, \boldsymbol{C}^{-1} \right).$$
(70)

For every $\boldsymbol{Q} \in ([T\mathcal{B}]_2^0, \text{sym})$, it holds that

$$\mathbb{I}^{\sharp*}: \boldsymbol{Q} = \boldsymbol{C}^{-1} \boldsymbol{Q} \boldsymbol{C}^{-1}.$$
(71)

In index notation, Equation (71) implies $(\mathbb{I}^{\sharp*} : \mathbf{Q})^{AB} =$ 866 $(C^{-1})^{AM}Q_{MN}(C^{-1})^{NB}$, which means that $\mathbb{I}^{\sharp*}$ raises the in-867 dices of Q through the inverse metric tensor C^{-1} rather than 868 through G^{-1} , the latter being the inverse of the metric tensor 869 *G* in the undeformed configuration. In analogy with \mathbb{K}^* and 870 \mathbb{M}^* , we also consider the fourth-order tensors 871

$$\mathbb{K}^{\sharp*} : ([T\mathcal{B}]_{2}^{0}, \text{sym}) \to ([T\mathcal{B}]_{0}^{2}, \text{sym}),$$

$$\mathbb{K}^{\sharp*} = \frac{1}{3}C^{-1} \otimes C^{-1},$$

$$\mathbb{M}^{\sharp*} : ([T\mathcal{B}]_{2}^{0}, \text{sym}) \to ([T\mathcal{B}]_{0}^{2}, \text{sym}),$$

$$\mathbb{M}^{\sharp*} = \mathbb{I}^{\sharp*} - \mathbb{K}^{\sharp*}.$$
(72b)

For every $\boldsymbol{Q} \in ([T\mathcal{B}]_2^0, \text{sym})$, we obtain

$$\mathbb{K}^{\sharp*}: \boldsymbol{O} = \frac{1}{2} \text{tr}(\boldsymbol{C}^{-1}\boldsymbol{O})\boldsymbol{C}^{-1}.$$
(73a)

$$\mathbb{M}^{\sharp*}: \boldsymbol{Q} = \boldsymbol{C}^{-1}\boldsymbol{Q}\boldsymbol{C}^{-1} - \frac{1}{3}\mathrm{tr}(\boldsymbol{C}^{-1}\boldsymbol{Q})\boldsymbol{C}^{-1}.$$
 (73b)

Note that the second-order tensor $\mathbb{M}^{\sharp *}$: Q is deviatoric in the sense that $tr[C(\mathbb{M}^{\sharp*} : Q)] = 0$. 874

Compliance with ethical standards

Conflict of interest: The authors declare that they have no 876 conflict of interest. 877

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