

Passive and active fibre reorientation in anisotropic materials.

Jacopo Ciambella^a, Giulio Lucci^b, Paola Nardinocchi^a, Luigi Preziosi^b

^a*Department of Structural and Geotechnical Engineering
Sapienza Università di Roma, I-00184 Rome, Italy*

^b*Department of Mathematical Sciences “G.L. Lagrange”
Politecnico di Torino, I-10129 Turin, Italy*

Abstract

We present a continuum model to describe the reorientation of an anisotropic material structure, characterized by two fiber families able to modify their orientations following different evolution dynamics. The evolution equations are derived in a thermodynamically consistent way, and passive and active contributions to the reorientation process are identified. It is shown that a weaker extension of a well-known coaxiality result holds. The transversely isotropic and orthotropic cases are then recovered by imposing the proper constraint on the fiber rotation. Applications to biological experiments on cell layers under stretch are discussed, [showing a good agreement between the model and the experimental results](#). Even though we focus on cell layers, our framework remains general and may be employed to describe reorientation in engineering materials.

Keywords: Fibre reorientation · Anisotropic materials · Nonlinear elasticity · Coaxiality · Cell orientation · Stress fibers

1. Introduction

2 The ability of actively changing the internal structure in response to ex-
3 ternal stimuli is a fundamental characteristic of biological tissues, cells, and

*Corresponding Author: G. Lucci, Department of Mathematical Sciences “G.L. Lagrange”, Politecnico di Torino, Corso Duca degli Abruzzi 24, I-10129 Turin, Italy. Tel: 0039 011 0907588. Email address: giulio.lucci@polito.it.

Email addresses: jacopo.ciambella@uniroma1.it (Jacopo Ciambella), giulio.lucci@polito.it (Giulio Lucci), paola.nardinocchi@uniroma1.it (Paola Nardinocchi), luigi.preziosi@polito.it (Luigi Preziosi)

1 organisms. Such a process, which is generally referred to as *remodeling*, may
2 be driven by growth, injuries, ageing or alterations in the chemo-mechanical
3 environment, to cite but a few examples [1, 2, 3, 4, 5]. As a consequence
4 of these prompts, a reorganization of the biological microstructure is often
5 observed, since the material constantly tries to adapt to the new conditions
6 in a dynamic way. Understanding and accurately describing the remodeling
7 of living structures is a major challenge in biomechanics, which has been of
8 great interest in recent years: from a mechanical viewpoint, several works
9 have focused on different features of remodeling, such as the coupling with
10 growth [4, 6, 7], the influence of diseases in shape alterations [8] or in ma-
11 terial properties, and the changes in the biological structure as plastic-like
12 irreversible distortions [9]; see also [10] and references therein for a detailed
13 overview of remodeling and growth in soft tissues.

14 Among the several types of remodeling that may be triggered by exter-
15 nal forces, reorientation is probably one of the most relevant as far as living
16 tissues and matter are concerned. For a biological material with a fiber-
17 reinforced structure, it can be defined as a change in the local orientation
18 of fibers due to some external actions, leading therefore to a reorganization
19 in the microstructure of the material. This phenomenon is frequently seen
20 in different situations: for instance, it is well known that the bones are able
21 to change the orientation of their internal fibers to functionally adapt to
22 environmental stimuli [11]. In particular, many biological materials with a
23 fiber-like microstructure are described as transversely isotropic, that is, a
24 single preferential direction exists and influences the mechanical behaviour:
25 significant examples are articular cartilage, tendons or skin, to cite but a
26 few [7, 12, 13, 14, 15, 16]. Instead, it is worth mentioning that tissues may
27 exhibit an orthotropic arrangement, which requires the description of two
28 distinct fiber directions. A significant example can be found in the circu-
29 latory system, where the myocardium and the arteries are known to have
30 three planes with distinct mechanical properties [17, 18, 19, 20, 21]. An-
31 other phenomenon related to orthotropy that has recently gathered some
32 attention is the cellular response to mechanical cues coming from the sur-
33 rounding microenvironment. In particular, experiments have demonstrated
34 that cells seeded on a two-dimensional substrate which is cyclically stretched
35 reorient their stress fibers and bodies to reach a stable configuration, charac-
36 terized by a precise angle between the cell axis and the direction of stretching
37 [22, 23, 24, 25, 26, 27, 28, 29].

38 A possible explanation of this behaviour, that relies on mechanical ar-
39 guments, has been firstly proposed by Livne et al. [22], who suggested that
40 cells attempt to minimize the elastic energy stored in the stretching process.

1 Recently, a generalization of the energy considered in [22] was presented in
2 [30, 31], where the system composed by the cells and the substrate was de-
3 scribed as an orthotropic material: the preferential directions were identified
4 with the cellular actin stress fibers and with the orthogonal protein network
5 interconnecting them [32, 33, 34, 35].

6 However, the nonlinear elastic description of cell orientations proposed in
7 [30] does not explicitly include a remodeling of the internal structure, even
8 if the material composed by substrate and cells is treated as orthotropic.
9 Instead, cells are viewed as passive fibers whose reorientation is dragged by
10 the deformation of the layer itself. A different point of view, even if limited
11 to transversely isotropic materials, has been recently proposed in [36, 37],
12 taking into account both passive and active changes in the fiber directions.
13 The introduction of a remodeling equation, that complements the usual
14 mechanical balances, allows to describe the evolution of fiber orientation
15 under mechanical [36] and magnetic [37] stimuli. Moreover, in [36], it was
16 shown that the stationary solutions of the remodeling equation are the ones
17 that make the remodeled stress and strain tensors coaxial [38].

18 It seems then quite natural to provide an extension of the results pre-
19 sented in [36] to a more general case, where an hyperelastic material is
20 endowed with a double fiber microstructure, as in the cellular example.
21 Furthermore, while in [30] the two fibers were fixed to be perpendicular,
22 it is of interest to characterize the remodeling of the different fiber families
23 independently, in a way that each fiber can change the orientation according
24 to its own rotation tensor.

25
26 In this paper, motivated by these observations, we propose a mechanical
27 model that describes the reorganization of an anisotropic material struc-
28 ture, characterized by two fiber families that can modify their orientation
29 following different evolution equations. Both active contributions affecting
30 the reorientation process, like external forces or stimuli, and purely passive
31 material remodeling can be incorporated in the proposed framework. In do-
32 ing so, we are able to extend an energetic modeling of cell alignment [22, 30]
33 using a fiber reorientation framework [36], in which the rotation of the cells
34 due to the mechanical prompt is treated as an additional variable for the
35 model. This allowed us to derive the reorientation equations in a rigorous
36 and thermodynamically consistent basis, recovering the results presented in
37 [30] as a particular case. At the same time, we provide an extension of the
38 fiber reorientation model proposed in [36], adapting it to anisotropic ma-
39 terials with two preferential directions that can change their orientation in
40 different manners, even though they can be properly coupled. The limit of

1 two families of fibers which cannot change their relative orientation, as done
2 for instance by Menzel [39] in the orthotropic case, is recovered within the
3 terms of a constrained model.

4 The paper is organized as follows. Section 2 presents the remodeling frame-
5 work in presence of a double-fibered anisotropic internal material structure,
6 through a generalization of the model proposed in [36]. Then, in Section
7 3, the stationary solutions of the reorientation equations for the fibers are
8 thoroughly studied, showing that a weaker extension of the coaxiality result
9 by Vianello [38] holds when the fiber reinforcements are able to reorient
10 without any constraint. Section 4 is devoted to the analysis of an in-plane
11 situation, where the fibers are supposed to lie in a plane during the whole
12 remodeling process. In this case, we are able to specify the general evolution
13 equations and the related stationary solutions, focusing for simplicity on a
14 minimal elastic energy which however allows relevant comparisons with ex-
15 perimental data. Section 5 is devoted to the discussion of a reduced model
16 based on an orthotropic material structure with fixed relative orientation
17 of the fibers, naturally emerging from the general framework, provided that
18 a constraint on fiber rotations is included. Applications of the proposed
19 models in the biological context of cell orientation are then presented in
20 Section 6. In particular we show that, as expected, the constrained model is
21 consistent with some already known results, such as the phenomenological
22 evolution equation for cell orientation postulated by Livne et al. [22] and
23 the stationary points of the energy derived in [30]. Moreover, the general
24 model is employed to describe stress fibers and cell nucleus reorientation,
25 showing a [good agreement](#) with experimental data by Roshanzadeh et al.
26 [29]. Finally, in Section 7 we provide some discussion and conclusions, as
27 well as future possible developments and applications of the present theory.
28 Appendix A is dedicated to the discussion of another possible choice for the
29 constraint between the fiber families.

30 **2. Remodeling framework**

31 Following the approach proposed in [36], we describe the reorientation
32 within the general framework of finite elasticity with remodeling presented
33 in [40]. We consider a material equipped with an anisotropic two families
34 of fibers (TFF) internal structure whose change of orientation is not simply
35 dragged by the deformation but described by additional state variables,
36 therefore including a remodeling of the fiber structure. The relationship
37 between fiber orientations and mechanical forces, as well as the constitutive

1 coupling between the two fiber families which characterize the anisotropic
 2 structure, are derived on a thermodynamically consistent basis.

3 2.1. Kinematics

4 Given a body, identified with a region \mathcal{B}_r of the Euclidean three-dimensional
 5 space \mathcal{E} , and a material point $X \in \mathcal{B}_r$, we consider the time-dependent
 6 map $\chi : \mathcal{B}_r \times \mathcal{T} \rightarrow \mathcal{E}$, called the deformation¹ of the body, that assigns
 7 to each point $X \in \mathcal{B}_r$ a point $x = \chi(X, t)$ at any instant t of the time
 8 interval \mathcal{T} and determines the current configuration $\mathcal{B} = \chi(\mathcal{B}_r, t)$ of the
 9 body at time t . As standard in finite elasticity, we introduce the displace-
 10 ment field $\mathbf{u} : \mathcal{B}_r \times \mathcal{T} \rightarrow \mathcal{V}$, with \mathcal{V} the translation space of \mathcal{E} , such that
 11 $x = \chi(X, t) = X + \mathbf{u}(X, t)$, and the fields

$$\mathbf{F}(X, t) = \nabla \chi(X, t) \quad \text{and} \quad \dot{\chi}(X, t) = \frac{\partial \chi}{\partial t}(X, t), \quad (2.1)$$

12 that represent the deformation gradient and the material velocity field, re-
 13 spectively. Clearly, $\mathbf{F} = \mathbf{I} + \nabla \mathbf{u}$, where \mathbf{I} is the identity tensor. We assume
 14 that at a certain time instant $t = t_0$, the body occupies the reference con-
 15 figuration, that is, $\chi(X, t_0) = X$ for each point in \mathcal{B}_r and $\mathbf{u}(X, t_0) = \mathbf{0}$.

16 In addition to the displacement field describing the current position of the
 17 body, we are interested in the evolution of the orientation of the anisotropic
 18 TFF internal structure, which in \mathcal{B}_r is based on the pair of material unit
 19 vector fields $\mathbf{a}_0 : \mathcal{B}_r \rightarrow \mathcal{V}$ and $\mathbf{b}_0 : \mathcal{B}_r \rightarrow \mathcal{V}$ (with $|\mathbf{a}_0| = |\mathbf{b}_0| = 1$); they
 20 represent the preferred directions that the internal structure endows to the
 21 material, i.e. the direction of each fiber at $X \in \mathcal{B}_r$. For practical usage, it
 22 is convenient to introduce the corresponding *structural or orientation ten-*
 23 *sors*² at \mathcal{B}_r , defined as $\mathbf{A}_0 = \mathbf{a}_0 \otimes \mathbf{a}_0$ and $\mathbf{B}_0 = \mathbf{b}_0 \otimes \mathbf{b}_0$. In the following,
 24 we will sometimes refer to the field \mathbf{a}_0 as the *primary structure* of the ma-
 25 terial, while \mathbf{b}_0 will be consequently called *secondary structure*. Both \mathbf{A}_0
 26 and \mathbf{B}_0 contribute to the material anisotropy and can reorient as prescribed
 27 by two different rotation fields, each one interpreted as the rotation of a
 28 single fiber family. In this respect, we will consider a time-dependent tensor
 29 field $\mathbf{R}_p : \mathcal{B}_r \times \mathcal{T} \rightarrow \mathbb{R}\text{ot}$ which represents the reorientation of the primary
 30 structure and changes the reference orientation from $\mathbf{a}_0(X)$ to

$$\mathbf{a}(X, t) = \mathbf{R}_p(X, t)\mathbf{a}_0(X),$$

¹We consider deformations that are twice continuously differentiable.

²Typically, they are denoted *structural tensors* in the mechanics of fiber-reinforced materials [18] and *orientation tensors* in the literature on nematic liquid crystals [41].

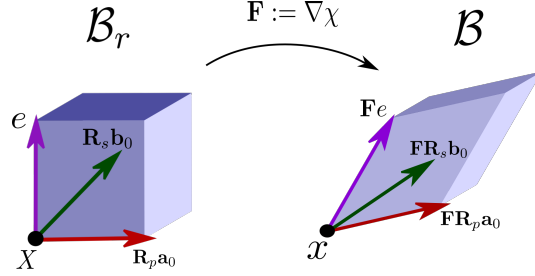


Figure 1: Schematic representation of the reorientation process at a material point. A material line element e in the reference configuration \mathcal{B}_r is deformed to the corresponding line element $\mathbf{F}e$ in the current configuration \mathcal{B} . Instead, the primary structure \mathbf{a}_0 firstly undergoes remodeling due to the rotation \mathbf{R}_p while the secondary structure \mathbf{b}_0 is reoriented according to another rotation \mathbf{R}_s .

1 or equivalently $\mathbf{A}_0(X)$ to

$$\mathbf{A}(X, t) = \mathbf{R}_p(X, t)\mathbf{A}_0(X)\mathbf{R}_p^T(X, t), \quad (2.2)$$

2 where $\mathbf{A} = \mathbf{a} \otimes \mathbf{a}$. Likewise, the second rotation field $\mathbf{R}_s : \mathcal{B}_r \times \mathcal{T} \rightarrow \mathbb{R}\text{ot}$
 3 describes the reorientation of the secondary structure and changes $\mathbf{b}_0(X)$ to

$$\mathbf{b}(X, t) = \mathbf{R}_s(X, t)\mathbf{b}_0(X)$$

4 and $\mathbf{B}_0(X)$ to

$$\mathbf{B}(X, t) = \mathbf{R}_s(X, t)\mathbf{B}_0(X)\mathbf{R}_s(X, t)^T, \quad (2.3)$$

5 having denoted $\mathbf{B} = \mathbf{b} \otimes \mathbf{b}$. Each fiber family of such an anisotropic material
 6 may change its orientation as time evolves; then, the deformation gradient
 7 tensor \mathbf{F} maps each remodeled fiber to the current configuration, as sketched
 8 in Fig. 1. With these definitions on hand, the state variables of the problem
 9 are the displacement and rotation fields, that is, the triple $(\mathbf{u}, \mathbf{R}_p, \mathbf{R}_s) \in$
 10 $\mathcal{V} \times \mathbb{R}\text{ot} \times \mathbb{R}\text{ot}$, and the corresponding velocity fields are identified by the
 11 time derivatives $(\dot{\mathbf{u}}, \dot{\mathbf{R}}_p\mathbf{R}_p^T, \dot{\mathbf{R}}_s\mathbf{R}_s^T) \in \mathcal{V} \times \mathbb{S}\text{kw} \times \mathbb{S}\text{kw}$, where $\mathbb{S}\text{kw}$ denotes
 12 the space of skew-symmetric tensors. Finally, we denote as $(\mathbf{w}, \mathbf{W}_p, \mathbf{W}_s) \in$
 13 $\mathcal{V} \times \mathbb{S}\text{kw} \times \mathbb{S}\text{kw}$ the associated virtual velocity fields.

14 *Remark.* Let us note that our kinematic framework does not filter out the
 15 rotation fields \mathbf{R}_p and \mathbf{R}_s such that $\mathbf{R}_p\mathbf{a}_0 = \mathbf{a}_0$ and $\mathbf{R}_s\mathbf{b}_0 = \mathbf{b}_0$, which are
 16 included in the admissible rotation fields even if they maintain the energy
 17 unchanged and so are of no interest in the remodeling problem.

1 *2.2. Balance equations*

2 The balance equations of the model are delivered by the principle of
 3 virtual working, based on the choice of the external and internal virtual
 4 workings defined as continuous, linear, real-valued functionals on the space
 5 of virtual velocities. By introducing forces and torques of the model which
 6 are working-conjugate to each kinematic variable, we write

$$\mathcal{W}_e(\mathbf{w}, \mathbf{W}_p, \mathbf{W}_s) = \int_{\mathcal{B}_r} (\mathbf{z} \cdot \mathbf{w} + \mathbf{Z}_p \cdot \mathbf{W}_p + \mathbf{Z}_s \cdot \mathbf{W}_s) dV + \int_{\partial_t \mathcal{B}_r} \mathbf{s} \cdot \mathbf{w} dA \quad (2.4)$$

7 for the external virtual working and

$$\mathcal{W}_i(\mathbf{w}, \mathbf{W}_p, \mathbf{W}_s) = \int_{\mathcal{B}_r} (\mathbf{S} \cdot \nabla \mathbf{w} + \boldsymbol{\Sigma}_p \cdot \mathbf{W}_p + \boldsymbol{\Sigma}_s \cdot \mathbf{W}_s) dV \quad (2.5)$$

8 for the internal virtual working. The pair (\mathbf{z}, \mathbf{s}) are forces per unit of (ref-
 9 erential) volume and area, respectively, while \mathbf{S} is the first Piola–Kirchhoff
 10 stress tensor. The pair of skew tensors $(\mathbf{Z}_i, \boldsymbol{\Sigma}_i)$, with $i = p, s$, are torques
 11 per unit of (referential) volume, and represent the outer and inner remodel-
 12 ing torques, as $\mathbf{W}_p, \mathbf{W}_s$ are skew-symmetric tensors. More specifically, the
 13 outer torques \mathbf{Z}_p and \mathbf{Z}_s represent external source terms which may affect
 14 fiber reorientation, for instance magnetic effects [37] or chemo-mechanical
 15 processes. On the other side, the inner remodeling torques $\boldsymbol{\Sigma}_p$ and $\boldsymbol{\Sigma}_s$ take
 16 into account the internal actions driving the reorientation of the primary
 17 and secondary material structures, respectively. It is worth remarking that,
 18 for what concerns the remodeling torques, our theory is of order zero-th,
 19 i.e., we do not take into account the gradients of the rotation fields, which
 20 instead may become relevant when the fibers are closely packed.

21 By enforcing the condition that the external and internal virtual working
 22 be equal for any virtual velocities $(\mathbf{w}, \mathbf{W}_p, \mathbf{W}_s) \in \mathcal{V} \times \text{Skw} \times \text{Skw}$ and for any
 23 subregion $\mathcal{R} \subset \mathcal{B}_r$, we obtain the following balance equations and associated
 24 boundary conditions:

$$\text{Div } \mathbf{S} + \mathbf{z} = \mathbf{0} \quad \text{and} \quad \boldsymbol{\Sigma}_p = \mathbf{Z}_p \quad \text{and} \quad \boldsymbol{\Sigma}_s = \mathbf{Z}_s \quad \text{in } \mathcal{B}_r, \quad (2.6)$$

$$\mathbf{u} = \hat{\mathbf{u}} \quad \text{on } \partial_u \mathcal{B}_r \quad \text{and} \quad \mathbf{S} \mathbf{m} = \mathbf{s} \quad \text{on } \partial_t \mathcal{B}_r, \quad (2.7)$$

26 with $\partial_u \mathcal{B}_r$ and $\partial_t \mathcal{B}_r$ denoting parts of the boundary $\partial \mathcal{B}_r$ where displacements
 27 and tractions are respectively prescribed, and \mathbf{m} denoting the unit normal
 28 to $\partial_t \mathcal{B}_r$. Equations (2.6) are the three balance equations of our theory. The
 29 first is the usual balance of mechanical forces, where inertial effects can be
 30 included in the bulk force \mathbf{z} , even if in the present model they are neglected

1 since the remodeling time scale is typically much longer than the inertial
 2 one. The second and third equations are the balances of the remodeling
 3 torques, driving the reorientation of the primary and secondary structures,
 4 respectively, once the appropriate constitutive equations will be considered.
 5 In particular, they prescribe that the outer and inner remodeling torques be
 6 equal. These equations generalise the theory presented in [36], where only
 7 a transversely isotropic internal structure was considered.
 8 The external working, that corresponds to balanced forces and torques and
 9 is evaluated on the actual velocity fields $(\dot{\mathbf{u}}, \dot{\mathbf{R}}_p \mathbf{R}_p^T, \dot{\mathbf{R}}_s \mathbf{R}_s^T)$, identifies the
 10 external actual power \mathcal{P}_e expended during the evolution of the continuum:

$$\begin{aligned}
 \mathcal{W}_e(\dot{\mathbf{u}}, \dot{\mathbf{R}}_p \mathbf{R}_p^T, \dot{\mathbf{R}}_s \mathbf{R}_s^T) &= \int_{\mathcal{B}_r} (\mathbf{z} \cdot \dot{\mathbf{u}} + \mathbf{Z}_p \cdot \dot{\mathbf{R}}_p \mathbf{R}_p^T + \mathbf{Z}_s \cdot \dot{\mathbf{R}}_s \mathbf{R}_s^T) dV + \int_{\partial_t \mathcal{B}_r} \mathbf{s} \cdot \dot{\mathbf{u}} dA \\
 &= \int_{\mathcal{B}_r} (\mathbf{S} \cdot \dot{\mathbf{F}} + \Sigma_p \cdot \dot{\mathbf{R}}_p \mathbf{R}_p^T + \Sigma_s \cdot \dot{\mathbf{R}}_s \mathbf{R}_s^T) = \mathcal{P}_e. \quad (2.8)
 \end{aligned}$$

11 2.3. Constitutive equations and energy imbalance

12 The constitutive recipes to describe the behaviour of the material with
 13 the reorientable anisotropic TFF internal structure are prescribed in a ther-
 14 modynamically consistent way, through the following steps. Firstly, we take
 15 into account a class of materials that admits an elastic strain energy density
 16 in the form

$$\varphi = \phi(\mathbf{E}, \mathbf{A}, \mathbf{B}) = \psi(\mathbf{E}, \mathbf{R}_p, \mathbf{R}_s), \quad (2.9)$$

17 dependent upon the deformation \mathbf{F} through the Green–Lagrange strain ten-
 18 sor $\mathbf{E} = \frac{1}{2}(\mathbf{F}^T \mathbf{F} - \mathbf{I})$, and on the remodeled orientation tensors \mathbf{A} and \mathbf{B}
 19 or, equivalently, on the rotations \mathbf{R}_p and \mathbf{R}_s that transform the primary
 20 and secondary structures. More specifically, following the theory of double-
 21 fibered finite elasticity, we assume that ϕ and ψ are isotropic functions of
 22 their arguments [42, 43].

23 We also assume that dissipation is only associated with the remodeling
 24 processes, and the dissipation density δ can be written as

$$\delta = \delta(\dot{\mathbf{R}}_p \mathbf{R}_p^T, \dot{\mathbf{R}}_s \mathbf{R}_s^T). \quad (2.10)$$

25 To derive constitutive equations that are consistent with the first and second
 26 laws of thermodynamics, we enforce the energy imbalance, stating that for
 27 any admissible process, characterised by the state variables $(\mathbf{u}, \mathbf{R}_p, \mathbf{R}_s)$, the
 28 time derivative of the energy must not exceed the external actual power

1 expended on the body along the same process, i.e., the dissipation must be
 2 positive:

$$\int_{\mathcal{B}_r} \delta dV = \mathcal{P}_e - \int_{\mathcal{B}_r} \dot{\varphi} dV \geq 0. \quad (2.11)$$

3 Within the framework discussed so far and using Eq. (2.9), we can write the
 4 time derivative of the elastic energy as

$$\dot{\varphi} = \frac{\partial \phi}{\partial \mathbf{E}} \cdot \dot{\mathbf{E}} + \frac{\partial \phi}{\partial \mathbf{A}} \cdot \dot{\mathbf{A}} + \frac{\partial \phi}{\partial \mathbf{B}} \cdot \dot{\mathbf{B}}. \quad (2.12)$$

At this point, it is useful to introduce the commutator operator between two tensors $[\cdot, \cdot] : \text{Lin} \times \text{Lin} \rightarrow \text{Lin}$, where Lin stands for the space of second-order tensors, defined as:

$$[\mathbf{X}, \mathbf{Y}] = \mathbf{X}\mathbf{Y} - \mathbf{Y}\mathbf{X}, \quad \forall \mathbf{X}, \mathbf{Y} \in \text{Lin}.$$

5 With this definition on hand, recalling Eqs. (2.2)-(2.3) we have

$$\dot{\mathbf{A}} = [\dot{\mathbf{R}}_p \mathbf{R}_p^T, \mathbf{A}] \quad \text{and} \quad \dot{\mathbf{B}} = [\dot{\mathbf{R}}_s \mathbf{R}_s^T, \mathbf{B}], \quad (2.13)$$

and, in general, for any $\mathbf{X}, \mathbf{Y} \in \text{Lin}$,

$$\mathbf{X} \cdot [\dot{\mathbf{R}}_p \mathbf{R}_p^T, \mathbf{A}] = \dot{\mathbf{R}}_p \mathbf{R}_p^T \cdot [\mathbf{X}, \mathbf{A}] \quad \text{and} \quad \mathbf{Y} \cdot [\dot{\mathbf{R}}_s \mathbf{R}_s^T, \mathbf{B}] = \dot{\mathbf{R}}_s \mathbf{R}_s^T \cdot [\mathbf{Y}, \mathbf{B}],$$

6 which allow Eq. (2.12) to be rewritten as

$$\dot{\varphi} = \frac{\partial \phi}{\partial \mathbf{E}} \cdot \dot{\mathbf{E}} + \left[\frac{\partial \phi}{\partial \mathbf{A}}, \mathbf{A} \right] \cdot \dot{\mathbf{R}}_p \mathbf{R}_p^T + \left[\frac{\partial \phi}{\partial \mathbf{B}}, \mathbf{B} \right] \cdot \dot{\mathbf{R}}_s \mathbf{R}_s^T. \quad (2.14)$$

7 The constitutive prescriptions for the stress \mathbf{S} and for the inner remodeling
 8 actions Σ_p and Σ_s have to satisfy the imbalance principle stated above,
 9 which can be rephrased in local form requiring that

$$\delta = \mathbf{S} \cdot \dot{\mathbf{F}} + \Sigma_p \cdot \dot{\mathbf{R}}_p \mathbf{R}_p^T + \Sigma_s \cdot \dot{\mathbf{R}}_s \mathbf{R}_s^T - \dot{\varphi} \geq 0 \quad (2.15)$$

10 has to be positive for every realizable process [44]. By using equations
 11 (2.12)-(2.14), we reduce the inequality (2.15) to:

$$\left(\mathbf{S} - \mathbf{F} \frac{\partial \phi}{\partial \mathbf{E}} \right) \cdot \dot{\mathbf{F}} + \left(\Sigma_p - \left[\frac{\partial \phi}{\partial \mathbf{A}}, \mathbf{A} \right] \right) \cdot \dot{\mathbf{R}}_p \mathbf{R}_p^T + \left(\Sigma_s - \left[\frac{\partial \phi}{\partial \mathbf{B}}, \mathbf{B} \right] \right) \cdot \dot{\mathbf{R}}_s \mathbf{R}_s^T \geq 0. \quad (2.16)$$

12 Since we assume that dissipation is only due to remodeling, the constitutive
 13 equation for the stress can be chosen as a standard hyperelastic law:

$$\mathbf{S} = \mathbf{F} \frac{\partial \phi}{\partial \mathbf{E}}, \quad \mathbf{S}_c = \mathbf{F}^{-1} \mathbf{S} = \frac{\partial \phi}{\partial \mathbf{E}}, \quad (2.17)$$

where \mathbf{S}_c is the second Piola-Kirchhoff stress tensor. The remainder of the dissipation inequality (2.16) can be satisfied on assuming that each inner remodeling torque can be decomposed into an elastic and a dissipative part:

$$\boldsymbol{\Sigma}_p = \boldsymbol{\Sigma}_p^{(e)} + \boldsymbol{\Sigma}_p^{(d)} = \left[\frac{\partial \phi}{\partial \mathbf{A}}, \mathbf{A} \right] + \mathbb{D}_p \dot{\mathbf{R}}_p \mathbf{R}_p^T, \quad (2.18)$$

$$\boldsymbol{\Sigma}_s = \boldsymbol{\Sigma}_s^{(e)} + \boldsymbol{\Sigma}_s^{(d)} = \left[\frac{\partial \phi}{\partial \mathbf{B}}, \mathbf{B} \right] + \mathbb{D}_s \dot{\mathbf{R}}_s \mathbf{R}_s^T, \quad (2.19)$$

- 1 where \mathbb{D}_p and \mathbb{D}_s are positive-definite fourth order tensors that represent the
 2 resistance to remodeling of the primary and secondary material structure,
 3 respectively.
 4 By substituting the constitutive equations (2.17)-(2.19) in the balance equa-
 5 tions (2.6), we are able to obtain two differential equations that describe the
 6 evolution of the rotation fields, and therefore of the associated fiber families.
 7 The general remodeling problem for the anisotropic TFF microstructure can
 8 then be expressed as the following system of nonlinear ordinary differential
 9 equations in $\mathcal{B}_r \times \mathcal{T}$:

$$\begin{cases} \mathbb{D}_p \dot{\mathbf{R}}_p \mathbf{R}_p^T = \mathbf{Z}_p - \left[\frac{\partial \phi}{\partial \mathbf{A}}, \mathbf{A} \right], \\ \mathbb{D}_s \dot{\mathbf{R}}_s \mathbf{R}_s^T = \mathbf{Z}_s - \left[\frac{\partial \phi}{\partial \mathbf{B}}, \mathbf{B} \right], \end{cases} \quad (2.20)$$

- 10 with initial conditions

$$\mathbf{R}_p = \mathbf{I} \quad \text{and} \quad \mathbf{R}_s = \mathbf{I} \quad \text{in} \quad \mathcal{B}_r \times \{t_0\}. \quad (2.21)$$

- 11 The problem (2.20) has to be solved together with the balance of forces (2.6)₁
 12 and the constitutive prescription (2.17). We observe that the elastic terms on
 13 the right-hand side of Eqs. (2.20), represented by the commutators, include
 14 the mutual interaction between the two fiber families, as it will be explicitly
 15 shown in the case of plane remodeling. Moreover, when $\mathbf{Z}_p = \mathbf{Z}_s = \mathbf{0}$, as we
 16 shall assume in the rest of the paper, the stationary solutions of the system
 17 (2.20) are determined by solving the equations

$$\left[\frac{\partial \phi}{\partial \mathbf{A}}, \mathbf{A} \right] = \mathbf{0} \quad \text{and} \quad \left[\frac{\partial \phi}{\partial \mathbf{B}}, \mathbf{B} \right] = \mathbf{0}. \quad (2.22)$$

18 2.4. Further constitutive prescriptions

- 19 For the calculations carried out in the following Sections, it is convenient
 20 to express the strain energy density in terms of the invariants of the tensors

1 \mathbf{E} , \mathbf{A} and \mathbf{B} . In doing so we introduce the following classical set of isotropic
 2 invariants of \mathbf{E} :

$$J_1 = \mathbf{E} \cdot \mathbf{I}, \quad J_2 = \mathbf{E}^\dagger \cdot \mathbf{I}, \quad J_3 = \det(\mathbf{I} + 2\mathbf{E}), \quad (2.23)$$

3 where $\mathbf{E}^\dagger = (\det \mathbf{E})\mathbf{E}^{-\text{T}}$, together with the anisotropic ones [45]

$$\begin{aligned} J_4 &= \mathbf{E} \cdot \mathbf{A}, & J_5 &= \mathbf{E}^2 \cdot \mathbf{A}, \\ J_6 &= \mathbf{E} \cdot \mathbf{B}, & J_7 &= \mathbf{E}^2 \cdot \mathbf{B}, \\ J_8 &= \mathbf{E} \cdot \text{sym}(\mathbf{A}\mathbf{B}) = (\mathbf{a} \cdot \mathbf{b})(\mathbf{E}\mathbf{a} \cdot \mathbf{b}). \end{aligned} \quad (2.24)$$

4 The anisotropic invariants J_4 and J_6 take into account the deformations
 5 along the directions of the primary and secondary internal structures, while
 6 J_5 and J_7 are also measures of fiber stretch but include the influence of shear
 7 on the fibers [46]; J_8 is instead used to account for the interactions between
 8 the two fiber families.³ If we use the representation theorem for isotropic
 9 scalar functions of three symmetric tensors (see [47, 48, 49]), we can write

$$\varphi = \phi(\mathbf{E}, \mathbf{A}, \mathbf{B}) = \hat{\phi}(J_1, J_2, J_3, J_4, J_5, J_6, J_7, J_8). \quad (2.25)$$

10 On adopting the representation (2.25) and denoting with $\hat{\phi}_i := \partial \hat{\phi} / \partial J_i$,
 11 $i = 1, \dots, 8$, one has:

$$\begin{aligned} \frac{\partial \phi}{\partial \mathbf{E}} &= \hat{\phi}_1 \mathbf{I} + \hat{\phi}_2 (J_1 \mathbf{I} - \mathbf{E}) + 2\hat{\phi}_3 J_3 (2\mathbf{E} + \mathbf{I})^{-1} + \hat{\phi}_4 \mathbf{A} + \hat{\phi}_5 (\mathbf{A}\mathbf{E} + \mathbf{E}\mathbf{A}), \\ &+ \hat{\phi}_6 \mathbf{B} + \hat{\phi}_7 (\mathbf{B}\mathbf{E} + \mathbf{E}\mathbf{B}) + \frac{1}{2} \hat{\phi}_8 (\mathbf{A}\mathbf{B} + \mathbf{B}\mathbf{A}), \end{aligned} \quad (2.26)$$

$$\frac{\partial \phi}{\partial \mathbf{A}} = \hat{\phi}_4 \mathbf{E} + \hat{\phi}_5 \mathbf{E}^2 + \frac{1}{2} \hat{\phi}_8 (\mathbf{E}\mathbf{B} + \mathbf{B}\mathbf{E}), \quad (2.27)$$

$$\frac{\partial \phi}{\partial \mathbf{B}} = \hat{\phi}_6 \mathbf{E} + \hat{\phi}_7 \mathbf{E}^2 + \frac{1}{2} \hat{\phi}_8 (\mathbf{E}\mathbf{A} + \mathbf{A}\mathbf{E}). \quad (2.28)$$

12 In the following, for the sake of a lighter notation, we will drop the hat from
 13 the derivatives of the strain energy with respect to the invariants.

³The coupling invariant J_8 may be defined in different ways. The definition of J_8 in (2.24) is the one proposed in [45]; however, there are other definitions that are sometimes used in the literature. In [18], for instance, the authors use $J_8 = \mathbf{a} \cdot \mathbf{E}\mathbf{b}$. In this paper we choose the former definition, which keeps the energy invariant under change of sign of either \mathbf{a} or \mathbf{b} .

1 *Remark.* It is often customary to express $\hat{\phi}$ in terms of the invariants of the
 2 right Cauchy–Green strain tensor $\mathbf{C} = \mathbf{F}^T \mathbf{F}$ rather than \mathbf{E} :

$$\begin{aligned}
 \mathbf{I}_1 &= \mathbf{C} \cdot \mathbf{I}, & \mathbf{I}_2 &= \mathbf{C}^\dagger \cdot \mathbf{I}, & \mathbf{I}_3 &= \det \mathbf{C}, \\
 \mathbf{I}_4 &= \mathbf{C} \cdot \mathbf{A}, & \mathbf{I}_5 &= \mathbf{C}^2 \cdot \mathbf{A}, \\
 \mathbf{I}_6 &= \mathbf{C} \cdot \mathbf{B}, & \mathbf{I}_7 &= \mathbf{C}^2 \cdot \mathbf{B}, \\
 \mathbf{I}_8 &= \mathbf{C} \cdot \text{sym}(\mathbf{A}\mathbf{B}).
 \end{aligned}
 \tag{2.29}$$

However, since the relationship $\mathbf{C} = 2\mathbf{E} + \mathbf{I}$ holds true, the two sets of invariants are connected by the following transformations:

$$\begin{aligned}
 \mathbf{J}_1 &= \frac{1}{2}(\mathbf{I}_1 - 3), & \mathbf{J}_2 &= \frac{1}{4}(\mathbf{I}_2 - 3) - \frac{1}{2}(\mathbf{I}_1 - 3), & \mathbf{J}_3 &= \mathbf{I}_3, \\
 \mathbf{J}_4 &= \frac{1}{2}(\mathbf{I}_4 - 1), & \mathbf{J}_5 &= \frac{1}{4}(\mathbf{I}_5 - 1) - \frac{1}{2}(\mathbf{I}_4 - 1), \\
 \mathbf{J}_6 &= \frac{1}{2}(\mathbf{I}_6 - 1), & \mathbf{J}_7 &= \frac{1}{4}(\mathbf{I}_7 - 1) - \frac{1}{2}(\mathbf{I}_6 - 1), \\
 \mathbf{J}_8 &= \frac{1}{2}(\mathbf{I}_8 - \mathbf{A} \cdot \mathbf{B}).
 \end{aligned}
 \tag{2.30}$$

3. Characterization of the remodeling stationary solutions

4 In [36], it was proved that, for materials equipped with a reorientable
 5 transversely isotropic internal structure described by a rotation field \mathbf{R} and
 6 in absence of external stimuli, the stationary solutions of the remodeling
 7 equations are those rotations which make stress \mathbf{S}_c and strain \mathbf{E} , or equiva-
 8 lently \mathbf{C} , coaxial. In addition, those rotations render the map

$$\sigma : \text{Rot} \ni \mathbf{R} \mapsto \sigma(\mathbf{R}) = \psi(\mathbf{C}, \mathbf{R})$$

9 stationary, where $\psi(\mathbf{C}, \mathbf{R})$ is the elastic strain energy (see Proposition 1 of
 10 [36] and also [38]).

11 In this Section, it is shown that these results can be partially extended to
 12 include the anisotropic double-fibered internal structure considered here. To
 13 do so, we first note that the results in [38], often referred to as *Vianello's*
 14 *coaxiality theorem*, hold true whatever class of material symmetry is con-
 15 sidered, either isotropic or orthotropic, granted that all fiber families are
 16 transformed under the same rotation acting on the body. It is then rea-
 17 sonable to expect that, under our more general framework in which two
 18 rotations appear, there could be a loss of equivalence between coaxiality
 19 and stationarity of the energy, as we shall prove in the following.

1 *3.1. Stationarity and coaxiality*

2 Before passing to the main result of this Section, we recall that two
 3 symmetric tensors \mathbf{U}, \mathbf{V} are said to be coaxial if they commute, or equiv-
 4 alently if their commutator vanishes, i.e. $[\mathbf{U}, \mathbf{V}] = \mathbf{0}$. Moreover, we prove
 5 the following relation:

$$[\frac{\partial\phi}{\partial\mathbf{A}}, \mathbf{A}] + [\frac{\partial\phi}{\partial\mathbf{B}}, \mathbf{B}] = [\mathbf{E}, \mathbf{S}_c]. \quad (3.31)$$

By recalling (2.27) and (2.28), the left-hand side of (3.31) can be written as

$$\begin{aligned} [\frac{\partial\phi}{\partial\mathbf{A}}, \mathbf{A}] + [\frac{\partial\phi}{\partial\mathbf{B}}, \mathbf{B}] &= \phi_4[\mathbf{E}, \mathbf{A}] + \phi_5[\mathbf{E}^2, \mathbf{A}] + \phi_6[\mathbf{E}, \mathbf{B}] + \phi_7[\mathbf{E}^2, \mathbf{B}] \\ &+ \frac{\phi_8}{2}(\mathbf{EBA} + \mathbf{BEA} - \mathbf{AEB} - \mathbf{ABE}) \\ &+ \frac{\phi_8}{2}(\mathbf{EAB} + \mathbf{AEB} - \mathbf{BEA} - \mathbf{BAE}) \\ &= \phi_4[\mathbf{E}, \mathbf{A}] + \phi_5[\mathbf{E}^2, \mathbf{A}] + \phi_6[\mathbf{E}, \mathbf{B}] + \phi_7[\mathbf{E}^2, \mathbf{B}] \\ &+ \frac{\phi_8}{2}\mathbf{E}(\mathbf{BA} + \mathbf{AB}) - \frac{\phi_8}{2}(\mathbf{AB} + \mathbf{BA})\mathbf{E} \\ &= \phi_4[\mathbf{E}, \mathbf{A}] + \phi_5[\mathbf{E}^2, \mathbf{A}] + \phi_6[\mathbf{E}, \mathbf{B}] + \phi_7[\mathbf{E}^2, \mathbf{B}] + \phi_8[\mathbf{E}, \text{sym}(\mathbf{AB})] \end{aligned}$$

which, in view of (2.26) and (2.17), is indeed equivalent to

$$\phi_4[\mathbf{E}, \mathbf{A}] + \phi_5[\mathbf{E}, \mathbf{AE} + \mathbf{EA}] + \phi_6[\mathbf{E}, \mathbf{B}] + \phi_7[\mathbf{E}, \mathbf{BE} + \mathbf{EB}] + \phi_8[\mathbf{E}, \text{sym}(\mathbf{AB})] = [\mathbf{E}, \mathbf{S}_c],$$

6 which proves Eq. (3.31). Then, at stationarity, since both the commutators
 7 on the left-hand side of (3.31) vanish, $[\mathbf{E}, \mathbf{S}_c] = \mathbf{0}$ holds, that is, the sta-
 8 tionary solutions of (2.20) in the passive case are rotations $(\mathbf{R}_p^*, \mathbf{R}_s^*)$ which
 9 make stress and strain coaxial. This result generalizes the one derived in
 10 [36] for transversely isotropic materials. We note however that the inverse
 11 statement is not necessarily true: in fact, coaxiality of stress and strain does
 12 not imply that both commutators have to vanish, and therefore need not be
 13 equivalent to a stationary solution of the remodeling system.

14 Taking into account this observations, we are now in the position of
 15 proving the following result ⁴.

⁴Proposition 1 is formulated in terms of the Green–Lagrange strain tensor \mathbf{E} . However, since the principal directions of \mathbf{E} and \mathbf{C} coincide, it could be equivalently stated in terms of the right Cauchy–Green strain tensor \mathbf{C} as was done in [36].

1 **Proposition 1.** *Let \mathbf{E} be a given deformation.*

(a) $(\mathbf{R}_p^*, \mathbf{R}_s^*)$ is a stationary solution of the passive remodeling system of equations if and only if $(\mathbf{R}_p^*, \mathbf{R}_s^*)$ is a critical point of the map

$$\sigma : \text{Rot} \times \text{Rot} \rightarrow \mathbb{R} \quad (3.32)$$

$$(\mathbf{R}_p, \mathbf{R}_s) \mapsto \sigma(\mathbf{R}_p, \mathbf{R}_s) = \phi(\mathbf{E}, \mathbf{A}, \mathbf{B}) = \phi(\mathbf{E}, \mathbf{R}_p \mathbf{A}_0 \mathbf{R}_p^T, \mathbf{R}_s \mathbf{B}_0 \mathbf{R}_s^T),$$

2 where ϕ is the strain energy density.

3 (b) If $(\mathbf{R}_p^*, \mathbf{R}_s^*)$ is a stationary solution of the remodelling system of equa-
 4 tions, then the stress $\mathbf{S}_c^* = \mathcal{S}(\mathbf{E}, \mathbf{R}_p^*, \mathbf{R}_s^*)$ and strain \mathbf{E} tensors are
 5 coaxial.

Proof. Statement (b) follows from the discussion carried out at the beginning of this Subsection. As a matter of fact, if $(\mathbf{R}_p^*, \mathbf{R}_s^*)$ is a stationary solution of the passive remodelling system (2.20), then

$$\left[\frac{\partial \phi}{\partial \mathbf{A}}, \mathbf{A} \right]^* = \mathbf{0} \quad \text{and} \quad \left[\frac{\partial \phi}{\partial \mathbf{B}}, \mathbf{B} \right]^* = \mathbf{0},$$

where we have used a superscript $*$ to denote quantities at stationarity. Consequently, by (3.31),

$$\mathbf{0} = \left[\frac{\partial \phi}{\partial \mathbf{A}}, \mathbf{A} \right]^* + \left[\frac{\partial \phi}{\partial \mathbf{B}}, \mathbf{B} \right]^* = [\mathbf{E}, \mathbf{S}_c^*]$$

6 and therefore the stress and strain tensors are coaxial. To prove statement
 7 (a), we follow the procedure put forward by Vianello [38]. In this case,
 8 however, we need to exploit the canonical isomorphism between the tangent
 9 space $\text{Rot}(\mathbf{R}_p, \mathbf{R}_s)$ to the product manifold $\text{Rot} \times \text{Rot}$ at $(\mathbf{R}_p, \mathbf{R}_s)$ and the
 10 product space $\text{Skw} \times \text{Skw}$, for which

$$\text{Rot}(\mathbf{R}_p, \mathbf{R}_s) = \{(\mathbf{W}_p \mathbf{R}_p, \mathbf{W}_s \mathbf{R}_s) \mid (\mathbf{W}_p, \mathbf{W}_s) \in \text{Skw} \times \text{Skw}\}.$$

In such a case, the derivative of the energy at $(\mathbf{R}_p, \mathbf{R}_s)$ in the direction $(\mathbf{W}_p \mathbf{R}_p, \mathbf{W}_s \mathbf{R}_s)$ becomes

$$\begin{aligned} \dot{\sigma}(\mathbf{R}_p, \mathbf{R}_s) &= D\sigma(\mathbf{R}_p, \mathbf{R}_s)[\mathbf{W}_p \mathbf{R}_p, \mathbf{W}_s \mathbf{R}_s] = \frac{\partial \phi}{\partial \mathbf{R}_p} \cdot \mathbf{W}_p \mathbf{R}_p + \frac{\partial \phi}{\partial \mathbf{R}_s} \cdot \mathbf{W}_s \mathbf{R}_s \\ &= \left[\frac{\partial \phi}{\partial \mathbf{A}}, \mathbf{A} \right] \cdot \mathbf{W}_p + \left[\frac{\partial \phi}{\partial \mathbf{B}}, \mathbf{B} \right] \cdot \mathbf{W}_s, \end{aligned}$$

1 which is null for every pair $(\mathbf{W}_p, \mathbf{W}_s) \in \text{Skw} \times \text{Skw}$ if and only if

$$2 \quad \left[\frac{\partial\phi}{\partial\mathbf{A}}, \mathbf{A}\right]^* = \mathbf{0} \quad \text{and} \quad \left[\frac{\partial\phi}{\partial\mathbf{B}}, \mathbf{B}\right]^* = \mathbf{0}, \quad (3.33)$$

3 i.e., if and only if $(\mathbf{R}_p^*, \mathbf{R}_s^*)$ is a stationary solution of the remodelling equa-
4 tions. \square

5 Previous derivations confirm the loss of equivalence between critical
6 points of the energy and coaxiality of stress and strain; indeed, in this
7 anisotropic TFF context, stationarity is a stronger requirement than coaxial-
8 ity, since it requires that both commutators have to vanish. In other words,
9 the set of stationary solutions is a proper subset of the set of rotations that
10 make the stress and strain coaxial.

10 4. In-plane remodeling

11 There are many situations of interest in which both fiber families are
12 constrained to rotate in the same plane; we refer to this situation as *in-plane*
13 *remodeling* and study it in more detail. By introducing an orthonormal
14 basis $\{\mathbf{e}_1, \mathbf{e}_2, \mathbf{e}_3\}$ in the vector space \mathcal{V} , we assume that fibers lie in the
15 plane $\{\mathbf{e}_1, \mathbf{e}_2\}$, whereas the rotations have an axis parallel to \mathbf{e}_3 ($\mathbf{R}_p \mathbf{e}_3 =$
16 $\mathbf{R}_s \mathbf{e}_3 = \mathbf{e}_3$). We also assume that the mobility tensors are both spherical and
17 determined by two positive scalar constants $m_p = \mu\tau_p$ and $m_s = \mu\tau_s$, with
18 $m_p \neq m_s$ in general, μ is a shear modulus and τ_p, τ_s are two characteristic
19 times of the remodeling processes:

$$\mathbb{D}_p = m_p \mathbb{I} \quad \text{and} \quad \mathbb{D}_s = m_s \mathbb{I}, \quad (4.34)$$

20 with \mathbb{I} the fourth-order identity tensor with components $\mathbb{I}_{ijkl} = \delta_{ik}\delta_{jl}$. Tak-
21 ing into account that $\dot{\mathbf{a}} = \dot{\mathbf{R}}_p \mathbf{R}_p^T \mathbf{a}$ and $\dot{\mathbf{b}} = \dot{\mathbf{R}}_s \mathbf{R}_s^T \mathbf{b}$, the remodeling system
22 of equations becomes

$$\begin{cases} m_p \dot{\mathbf{a}} = -\left[\frac{\partial\phi}{\partial\mathbf{A}}, \mathbf{A}\right] \mathbf{a}, \\ m_s \dot{\mathbf{b}} = -\left[\frac{\partial\phi}{\partial\mathbf{B}}, \mathbf{B}\right] \mathbf{b}. \end{cases} \quad (4.35)$$

23 On recalling Eq. (2.27)-(2.28), the previous system can be recast in the
24 following form:

$$\begin{cases} m_p \dot{\mathbf{a}} = (\mathbf{A} - \mathbf{I})(\phi_4 \mathbf{Ea} + \phi_5 \mathbf{E}^2 \mathbf{a} + \frac{\phi_8}{2} \mathbf{BEa} + \frac{\phi_8}{2} \mathbf{EBa}), \\ m_s \dot{\mathbf{b}} = (\mathbf{B} - \mathbf{I})(\phi_6 \mathbf{Eb} + \phi_7 \mathbf{E}^2 \mathbf{b} + \frac{\phi_8}{2} \mathbf{AEb} + \frac{\phi_8}{2} \mathbf{EAb}). \end{cases} \quad (4.36)$$

1 Since the problem is in-plane, it is convenient to introduce a parametrization
 2 in terms of the remodeling angles θ_p and θ_s , that is, we set $\mathbf{a} = \cos \theta_p \mathbf{e}_1 + \sin \theta_p \mathbf{e}_2$,
 3 $\mathbf{b} = \cos \theta_s \mathbf{e}_1 + \sin \theta_s \mathbf{e}_2$. In addition, without loss of generality, we can as-
 4 sume that \mathbf{e}_1 , \mathbf{e}_2 and \mathbf{e}_3 are the principal strain directions of \mathbf{E} associated
 5 with the principal strains ε_1 , ε_2 and ε_3 .
 6 Equations (4.36) correspond therefore to the following system of scalar evo-
 7 lution equations for the angles:

$$\begin{cases} 2 m_p \dot{\theta}_p = [\phi_4 + \frac{1}{2}\phi_8 + \phi_5(\varepsilon_1 + \varepsilon_2)] (\varepsilon_1 - \varepsilon_2) \sin 2\theta_p + \frac{1}{2}\phi_8(\varepsilon_1 + \varepsilon_2) \sin [2(\theta_p - \theta_s)], \\ 2 m_s \dot{\theta}_s = [\phi_6 + \frac{1}{2}\phi_8 + \phi_7(\varepsilon_1 + \varepsilon_2)] (\varepsilon_1 - \varepsilon_2) \sin 2\theta_s + \frac{1}{2}\phi_8(\varepsilon_1 + \varepsilon_2) \sin [2(\theta_s - \theta_p)], \end{cases} \quad (4.37)$$

8 to be solved with the initial conditions $\theta_p(0) = \theta_{p0}$ and $\theta_s(0) = \theta_{s0}$, in which
 9 θ_{p0} , θ_{s0} are the referential primary and secondary orientation angles.

10 The following observations are duly noted:

- 11 1. depending on the representation form of the elastic strain energy, the
 12 strain component ε_3 of \mathbf{E} may or may not enter the equations (4.37);
 13 indeed, ε_3 does not appear explicitly, but it may be included in one of
 14 the energy derivatives ϕ_i , $i = 4 \dots 8$. For the sake of simplicity, in the
 15 following we will assume $\varepsilon_3 = 0$;
- 16 2. the directions of principal strain \mathbf{e}_1 , \mathbf{e}_2 , that correspond to $\theta_p = k\frac{\pi}{2}$ and
 17 $\theta_s = j\frac{\pi}{2}$ ($k, j = 0, 1, 2, \dots$) are stationary solutions of the remodelling
 18 equations (4.37);
- 19 3. if $\varepsilon_1 = \varepsilon_2 = \varepsilon$, i.e., the deformation is equibiaxial, the system of
 20 equations (4.37) simplifies into

$$\begin{cases} m_p \dot{\theta}_p = \frac{1}{2}\phi_8 \varepsilon \sin [2(\theta_p - \theta_s)] \\ m_s \dot{\theta}_s = \frac{1}{2}\phi_8 \varepsilon \sin [2(\theta_s - \theta_p)] \end{cases} \quad (4.38)$$

21 and the stationary solutions are achieved either if $\phi_8(\varepsilon, \theta_p, \theta_s) = 0$ or if
 22 $\sin 2(\theta_p - \theta_s) = 0$. The latter condition means that $\Delta\theta = \theta_s - \theta_p = k\frac{\pi}{2}$
 23 ($k = 0, 1, \dots$), i.e., the fibers become either parallel or orthogonal, yet
 24 their “absolute” angle with respect to \mathbf{e}_1 remains undetermined. On
 25 the other hand, if $\phi_8 = 0$ identically, meaning that the energy does not
 26 depend on J_8 , then no remodeling occurs, as found also for transversely
 27 isotropic materials [36].

28 To solve the remodeling problem (4.37) and make comparisons with ex-
 29 periments, an *ansatz* on the strain energy function shall be made. In this

1 respect, one could consider the most general form of the strain energy func-
 2 tion which is quadratic in the deformation measures. To do so, we generalize
 3 the well-known Saint Venant–Kirchhoff model, that in its original formula-
 4 tion only depends on the isotropic invariants, to a double-fibered material
 5 [50, 51]. The minimal representation of that energy which provides the sim-
 6 plest coupling between the fibers and allows to fit data from experiments on
 7 biaxial tests is:

$$\begin{aligned} \hat{\phi}(J_1, \dots, J_8) = & \frac{1}{2}k_1 J_1^2 + k_2 J_2 + \frac{1}{2}k_4 J_4^2 + \frac{1}{2}k_6 J_6^2 + \\ & + k_{14} J_1 J_4 + k_{16} J_1 J_6 + \frac{1}{2} k_8 J_8^2. \end{aligned} \quad (4.39)$$

With these assumptions, the remodelling problem takes the following form:

$$2 m_p \dot{\theta}_p = (1+r)\varepsilon_1^2 F(\theta_p, r; k_4, k_{14}) + \frac{1}{2}\varepsilon_1^2 k_8 [G(\theta_p, \theta_s, r) + H(\theta_p, \theta_s, r)] , \quad (4.40)$$

$$2 m_s \dot{\theta}_s = (1+r)\varepsilon_1^2 F(\theta_s, r; k_6, k_{16}) + \frac{1}{2}\varepsilon_1^2 k_8 [G(\theta_s, \theta_p, r) + H(\theta_s, \theta_p, r)] , \quad (4.41)$$

where we have introduced the biaxiality ratio $r := -\varepsilon_2/\varepsilon_1$ between the transverse and longitudinal deformation, while the functions F , G and H are defined as

$$F(\theta_p, r; k_4, k_{14}) := (k_4(\cos^2 \theta_p - r \sin^2 \theta_p) + k_{14}(1-r)) \sin 2\theta_p , \quad (4.42)$$

$$G(\theta_p, \theta_s, r) := 2(1-r) \cos^2(\theta_p - \theta_s) (\cos \theta_p \cos \theta_s - r \sin \theta_p \sin \theta_s) \sin(\theta_p - \theta_s) , \quad (4.43)$$

$$H(\theta_p, \theta_s, r) := (1+r) \cos(\theta_p - \theta_s) (\cos \theta_p \cos \theta_s - r \sin \theta_p \sin \theta_s) \sin 2\theta_p . \quad (4.44)$$

8 A discussion of the reorientation dynamics prescribed by the equations above
 9 is postponed to Section 6, where inspired by experiments on cell layers we
 10 will be able to estimate some of the constitutive coefficients $(k_4, k_{14}, k_6, k_{16}, k_8)$.
 11

12 *Remark.* The representation form of the equations (4.40) and (4.41) suggests
 13 to identify a system of elastic actions which contribute to the time rate of
 14 the fiber orientation angles θ_p and θ_s . Firstly, we introduce the pair

$$\tau_p := (1+r)\varepsilon_1^2 F(\theta_p, r; k_4, k_{14}) \quad \text{and} \quad \tau_s := (1+r)\varepsilon_1^2 F(\theta_s, r; k_6, k_{16}) . \quad (4.45)$$

1 Then, we note that

$$G(\theta_s, \theta_p, r) = -G(\theta_p, \theta_s, r). \quad (4.46)$$

2 As a consequence, we can identify the following mutual elastic interaction
3 between the fibers:

$$\tau_{ps} := \frac{1}{2} \varepsilon_1^2 k_8 G(\theta_p, \theta_s, r), \quad (4.47)$$

4 dependent upon the constitutive coefficient k_8 . In addition, we can define

$$\tau_{Hp} := \frac{1}{2} \varepsilon_1^2 k_8 H(\theta_p, \theta_s, r) \quad \text{and} \quad \tau_{Hs} := \frac{1}{2} \varepsilon_1^2 k_8 H(\theta_s, \theta_p, r). \quad (4.48)$$

5 With this on hand, the remodeling equations are restated as

$$2m_p \dot{\theta}_p = \tau_p + \tau_{ps} + \tau_{Hp} \quad \text{and} \quad 2m_s \dot{\theta}_s = \tau_s - \tau_{ps} + \tau_{Hs}, \quad (4.49)$$

6 which have the following physical interpretation: the time evolution of the
7 orientation of the primary (resp. secondary) angle depends on the elastic
8 actions τ_p (resp. τ_s), τ_{Hp} (resp. τ_{Hs}) and τ_{ps} (resp. $-\tau_{ps}$). τ_p is defined in
9 Eq. (4.45) and depends on the orientation θ_p (resp. θ_s) of the fiber itself
10 and on the stretches $\varepsilon_1, \varepsilon_2$. On the other hand, the interactions τ_{ps} and τ_{Hp}
11 (resp. τ_{Hs}) depend on both the orientations θ_p and θ_s and vanish when the
12 two fiber families are uncoupled, that is when k_8 is zero.

13 It is of particular interest the evaluation of $\tau_p, \tau_s, \tau_{Hp}, \tau_{Hs}$ and τ_{ps} when
14 the stretch is equibiaxial, i.e., for $r = -1$, and then $\varepsilon_1 = \varepsilon_2 = \varepsilon$. In such a
15 case, Eqs. (4.43), (4.47) and (4.48) imply $\tau_p = \tau_s = \tau_{Hp} = \tau_{Hs} = 0$ and

$$\tau_{ps} = 2k_8 \varepsilon^2 \cos^3(\theta_p - \theta_s) \sin(\theta_p - \theta_s), \quad (4.50)$$

16 meaning that the only non-zero term in the right-hand side of (4.49) is τ_{ps} .
17 Since stationarity under equibiaxial stretch is attained if $\sin 2(\theta_p - \theta_s) = 0$,
18 Eq. (4.50) states that the stationary value of the elastic interaction is zero.

19 5. Constrained reorientation

20 There may be practical cases in which the rotation of a fiber family
21 induces the same rotation in the other family, in the sense that, during
22 the remodeling process, the difference between the reoriented fiber angles
23 remains the same. This behaviour is introduced in the modeling framework
24 detailed above by properly constraining the elastic energy with a Lagrange
25 multiplier. Among the many possible choices (see for instance [52]), guided

1 by the experiments in [22] in which the *relative orientation* is fixed, we
 2 require that the primary \mathbf{R}_p and secondary \mathbf{R}_s rotation tensors are equal
 3 at any time instant. As such, we introduce the representation of rotation
 4 tensors through the exponential maps $\mathcal{B}_r \times \mathcal{T} \rightarrow \text{Skw}$ [52] given by

$$\mathbf{R}_p = \exp \boldsymbol{\Omega}_p \quad \text{and} \quad \mathbf{R}_s = \exp \boldsymbol{\Omega}_s. \quad (5.51)$$

5 Then, $\boldsymbol{\Omega}_p = \boldsymbol{\Omega}_s$ implies $\mathbf{R}_p = \mathbf{R}_s$, and the constraint expression takes the
 6 form

$$\mathbf{c}(\boldsymbol{\Omega}_p, \boldsymbol{\Omega}_s) = \boldsymbol{\Omega}_p - \boldsymbol{\Omega}_s, \quad (5.52)$$

7 with the corresponding constrained energy given by

$$\tilde{\varphi}(\mathbf{E}, \mathbf{A}, \mathbf{B}) = \phi(\mathbf{E}, \mathbf{A}, \mathbf{B}) + \mathbf{K} \cdot \mathbf{c}(\boldsymbol{\Omega}_p, \boldsymbol{\Omega}_s). \quad (5.53)$$

8 Hence, the skew tensor field \mathbf{K} is the Lagrange multiplier, which physically
 9 represents the reaction needed to maintain the relative rotation between the
 10 two fiber families the same. Equation (5.52) implies

$$\dot{\mathbf{c}}(\mathbf{R}_p, \mathbf{R}_s) = \dot{\mathbf{R}}_p \mathbf{R}_p^T - \dot{\mathbf{R}}_s \mathbf{R}_s^T. \quad (5.54)$$

On substituting (5.52) and (5.53) into the dissipation inequality, we arrive
 at

$$\begin{aligned} & \left(-\mathbf{S} + \mathbf{F} \frac{\partial \phi}{\partial \mathbf{E}} \right) \cdot \dot{\mathbf{F}} + \dot{\mathbf{K}} \cdot \mathbf{c}(\boldsymbol{\Omega}_p, \boldsymbol{\Omega}_s) \\ & + \left(-\boldsymbol{\Sigma}_p + \left[\frac{\partial \phi}{\partial \mathbf{A}}, \mathbf{A} \right] + \mathbf{K} \right) \cdot \dot{\mathbf{R}}_p \mathbf{R}_p^T \\ & + \left(-\boldsymbol{\Sigma}_s + \left[\frac{\partial \phi}{\partial \mathbf{B}}, \mathbf{B} \right] - \mathbf{K} \right) \cdot \dot{\mathbf{R}}_s \mathbf{R}_s^T \leq 0, \end{aligned}$$

11 which leads to the following system of *constrained* remodelling equations:

$$\begin{cases} \mathbb{D}_p \dot{\mathbf{R}}_p \mathbf{R}_p^T = -\left[\frac{\partial \phi}{\partial \mathbf{A}}, \mathbf{A} \right] - \mathbf{K}, \\ \mathbb{D}_s \dot{\mathbf{R}}_s \mathbf{R}_s^T = -\left[\frac{\partial \phi}{\partial \mathbf{B}}, \mathbf{B} \right] + \mathbf{K}, \\ \mathbf{c}(\boldsymbol{\Omega}_p, \boldsymbol{\Omega}_s) = \mathbf{0}. \end{cases} \quad (5.55)$$

12 Summing up the first two equations in (5.55), we obtain an evolution equa-
 13 tion independent of the Lagrange multiplier \mathbf{K} :

$$\mathbb{D}_p \dot{\mathbf{R}}_p \mathbf{R}_p^T + \mathbb{D}_s \dot{\mathbf{R}}_s \mathbf{R}_s^T = -\left[\frac{\partial \phi}{\partial \mathbf{A}}, \mathbf{A} \right] - \left[\frac{\partial \phi}{\partial \mathbf{B}}, \mathbf{B} \right]$$

1 which, at stationarity, gives

$$[\frac{\partial \phi}{\partial \mathbf{A}}, \mathbf{A}]^* + [\frac{\partial \phi}{\partial \mathbf{B}}, \mathbf{B}]^* = \mathbf{0} = [\mathbf{E}, \mathbf{S}_c]^*, \quad (5.56)$$

2 recovering the result in Eq. (3.31). On the other hand, by subtracting (5.55)₁
3 from (5.55)₂ and using (5.56), at stationarity we obtain

$$\mathbf{K}^* = [\frac{\partial \phi}{\partial \mathbf{B}}, \mathbf{B}]^*. \quad (5.57)$$

4 which furnishes the stationary values of the Lagrange multiplier \mathbf{K}^* . Equa-
5 tions (5.56), (5.57) together with the constraint equation (5.55)₃ are a sys-
6 tem of nine equations. The nine unknowns are the six components (a_i, b_i)
7 ($i = 1, 2, 3$) and the three independent components of $\mathbf{K} \in \text{Skw}$. Therefore,
8 in this constrained case, the equivalence between coaxiality of stress and
9 strain and stationarity of the solutions of the remodeling equation is recov-
10 ered thanks to the introduction of the multiplier \mathbf{K} . Indeed, if $(\mathbf{R}_p^*, \mathbf{R}_s^*)$
11 is a pair of rotations compatible with the constraint that makes stress and
12 strain coaxial, then (5.56) is trivially satisfied. However, the reaction term
13 \mathbf{K} and the constraint equation guarantee that $(\mathbf{R}_p^*, \mathbf{R}_s^*)$ is also a stationary
14 solution, as it is possible to find a value of \mathbf{K}^* satisfying Eq. (5.57) once \mathbf{a}^*
15 and \mathbf{b}^* are known.

16 5.1. In-plane remodeling under constraint

17 When in-plane remodeling is considered, we set $\boldsymbol{\Omega}_p = (\theta_p - \theta_{p0}) \star \mathbf{e}_3$ and
18 $\boldsymbol{\Omega}_s = (\theta_s - \theta_{s0}) \star \mathbf{e}_3$, where the operator $\star : \mathcal{V} \rightarrow \text{Skw}$ associates to each
19 vector \mathbf{w} a skew tensor \mathbf{W} whose axial vector is \mathbf{w} , i.e. $(\star \mathbf{w})\mathbf{u} = \mathbf{w} \times \mathbf{u}$ for
20 any $\mathbf{u} \in \mathcal{V}$. With this notation, we have $\mathbf{K} = \kappa \star \mathbf{e}_3$. As a consequence, the
21 constraint reduces to

$$c(\theta_p, \theta_s) = (\theta_p - \theta_s) - (\theta_{p0} - \theta_{s0}), \quad (5.58)$$

whence the remodeling equations become

$$\begin{cases} 2 m_p \dot{\theta}_p = (1+r) \varepsilon_1^2 F(\theta_p, r; k_4, k_{14}) + \frac{1}{2} \varepsilon_1^2 k_8 [G(\theta_p, \theta_s, r) + H(\theta_p, \theta_s, r)] - 2\kappa \\ 2 m_s \dot{\theta}_s = (1+r) \varepsilon_1^2 F(\theta_s, r; k_6, k_{16}) + \frac{1}{2} \varepsilon_1^2 k_8 [G(\theta_s, \theta_p, r) + H(\theta_s, \theta_p, r)] + 2\kappa \\ \theta_p - \theta_s = \theta_{p0} - \theta_{s0}. \end{cases} \quad (5.59)$$

22 The constraint equation (5.59)₃ implies

$$\dot{\theta}_p - \dot{\theta}_s = 0$$

from which by summing and subtracting (5.59)₁ and (5.59)₂ we get

$$2(m_p + m_s)\dot{\theta}_p = (1+r)\varepsilon_1^2 [F(\theta_p, r; k_4, k_{14}) + F(\theta_s, r; k_6, k_{16})] + \frac{1}{2}\varepsilon_1^2 k_8 [H(\theta_p, \theta_s, r) + H(\theta_s, \theta_p, r)] \quad (5.60)$$

and

$$2(m_p - m_s)\dot{\theta}_p = (1+r)\varepsilon_1^2 [F(\theta_p, r; k_4, k_{14}) - F(\theta_s, r; k_6, k_{16})] + \frac{1}{2}\varepsilon_1^2 k_8 [2G(\theta_p, \theta_s, r) + H(\theta_p, \theta_s, r) - H(\theta_s, \theta_p, r)] - 4\kappa, \quad (5.61)$$

1 respectively, recalling (4.46). When the deformation is equibiaxial, $r = -1$
 2 and by using the definitions of the functions F , G and H , Eq. (5.60) gives

$$2(m_p + m_s)\dot{\theta}_p = 0, \quad (5.62)$$

3 thus $\dot{\theta}_s = \dot{\theta}_p = 0$ and no evolution occurs as happened in the transversely
 4 isotropic case [36]. Moreover, Eq. (5.61) gives

$$\kappa = k_8 \varepsilon_1^2 \cos^3(\theta_{p0} - \theta_{s0}) \sin(\theta_{p0} - \theta_{s0}) \quad (5.63)$$

5 and shows that the reaction κ is zero either for fibre initially parallel, i.e.,
 6 $\theta_{p0} - \theta_{s0} = k\pi$, or orthogonal, i.e., $\theta_{p0} - \theta_{s0} = \frac{\pi}{2} + k\pi$ ($k = 0, 1, \dots$).

7 *Remark.* Other choices for the constraint (5.52) are indeed possible. One
 8 based on constraining the squared inner product between fiber vectors is
 9 presented and discussed in the Appendix.

10 6. Examples from Biology

11 In this Section, we illustrate some applications of our framework in the
 12 field of biology, which allow us to discuss interesting implications of the
 13 model and to make a comparison with experimental data. In particular, we
 14 focus on describing cell orientation under stretch: as mentioned in the In-
 15 troduction, when a monolayer of cells is stretched biaxially, a reorientation
 16 happens until each cell finds a stable configuration, characterized by a cer-
 17 tain angle with respect to the direction of greatest stretch. For more details
 18 on the biology and mechanics of this phenomenon, we refer the reader to
 19 [22, 27, 28, 30, 53] and references therein.

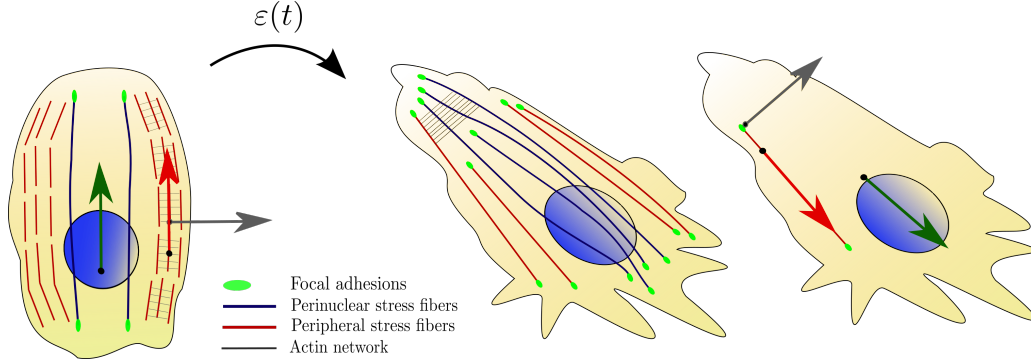


Figure 2: Sketch of the cell structure and reorientation process. For comparison with experimental assays in [22], following [30], we consider the reorientation of the actin peripheral stress fibers (in red) which determine the alignment of the cell body, while the lateral protein network (in gray) is constrained to remain orthogonal to the SF. Instead, to compare our model with results in [29], we consider the peripheral stress fibers as the primary structure, whose orientation is changed according to \mathbf{R}_p , while the nucleus and the perinuclear stress fibers (in blue) that drive its orientation represent the secondary structure, evolving with \mathbf{R}_s .

1 6.1. The case of constrained reorientation

2 Livne and coworkers [22] firstly proposed a model of cell orientation
3 based on the minimization of the stored elastic energy with respect to the
4 angle between the cell and the stretching direction. Later, a generalization
5 of their results using a nonlinear orthotropic energy was presented in [30],
6 with the two fiber families representing the cell aligned stress fibers and
7 the orthogonal linking protein network, as sketched in Figure 2. In the
8 following, we show that our model is able to recover the results in [22, 30]
9 as a particular case, without the need of any phenomenological justification
10 of the evolution equation.

11 As a starting point, following [30], we consider the cell monolayer as an
12 hyperelastic anisotropic material, characterized by the strain energy function
13 defined in Eq. (4.39). Then, we introduce the two assumptions also (implicit-
14 ly) done in [22]: (1) the families of fibers are orthogonal in the reference
15 configuration and constrained to remain orthogonal during remodeling; and
16 (2) the fibers lie in the plane of unit normal \mathbf{e}_3 . Finally, we consider a
17 deformation of the substrate defined by $\mathbf{E} = \text{diag}(\varepsilon_1, -r\varepsilon_1, 0)$.

18 In doing so, we are able to apply the constrained in-plane remodeling
19 theory derived in Section 5.1 and to write the reorientation equations using
20 (5.59). Taking into account that $\theta_{p0} - \theta_{s0} = \pi/2$, the constraint becomes
21 $c(\theta_p, \theta_s) = (\theta_p - \theta_s) - \pi/2 = 0$, which also implies $\dot{c}(\theta_p, \theta_s) = \dot{\theta}_p - \dot{\theta}_s = 0$.

1 Moreover, the orthogonality of the fibers yields

$$G(\theta_p, \theta_s, r) = G(\theta_s, \theta_p, r) = 0 \quad \text{and} \quad H(\theta_p, \theta_s, r) = H(\theta_s, \theta_p, r) = 0,$$

2 and the remodeling equations are simplified as

$$\begin{cases} 2m_p \dot{\theta}_p = (1+r)\varepsilon_1^2 F(\theta_p, r; k_4, k_{14}) - 2\kappa, \\ 2m_s \dot{\theta}_s = (1+r)\varepsilon_1^2 F(\theta_s, r; k_6, k_{16}) + 2\kappa. \end{cases} \quad (6.64)$$

3 Summing up the two equations and exploiting the constraint, we can
4 obtain an evolution equation for the cell orientation θ_p :

$$\widehat{m} \dot{\theta}_p = (1+r)\varepsilon_1^2 \left[\widehat{k}_p (\cos^2 \theta_p - r \sin^2 \theta_p) + k_m (1-r) \right] \sin 2\theta_p, \quad (6.65)$$

5 where

$$\widehat{m} := 2(m_p + m_s), \quad \widehat{k}_p := k_4 + k_6, \quad k_m := k_{14} - k_{16} - k_6.$$

6 The stationary solutions of Eq. (6.65) can be readily identified: they corre-
7 spond either to the cell being aligned with the principal directions of strain,
8 i.e. $\theta_p^* = k\pi/2, k \in \mathbb{Z}$, or to the oblique orientations defined by

$$\cos^2 \theta_p^* = 1 + \frac{k_m}{\widehat{k}_p} - \frac{1}{1+r} \left(1 + 2\frac{k_m}{\widehat{k}_p} \right) = \frac{1}{2} + \mathcal{K} \left(\frac{1}{2} - \frac{1}{1+r} \right), \quad (6.66)$$

9 where $\mathcal{K} := 1 + 2k_m/\widehat{k}_p$. Such a result is coherent with previous findings
10 characterizing the preferential orientations of cells on a stretched substrate,
11 both in linear elasticity [22], nonlinear elasticity [30] and viscoelasticity [31].
12 However, in our model we did not postulate the evolution equation (6.65)
13 as done in previous works, but rather derived it from a more general frame-
14 work lying on balance principles and thermodynamics. Clearly, the oblique
15 stationary solutions only exist when $r \neq -1$ and the right-hand side of
16 Eq. (6.66) has a value between 0 and 1; for details on the bifurcation anal-
17 ysis of the orientations, see [30].

18 The Lagrange multiplier κ , that represents the reaction needed to keep
19 the fibers orthogonal, can be evaluated at stationarity through

$$4\kappa^* = (1+r)\varepsilon_1^2 \left[(k_4 - k_6)(\cos^2 \theta_p^* - r \sin^2 \theta_p^*) + (k_{14} + k_{16} + k_6)(1-r) \right] \sin 2\theta_p^*. \quad (6.67)$$

20 We observe that, when the cell is aligned with the principal strain directions,
21 $\kappa^* = 0$. Indeed the same holds true when $r = -1$ (see (6.65)), i.e. in the

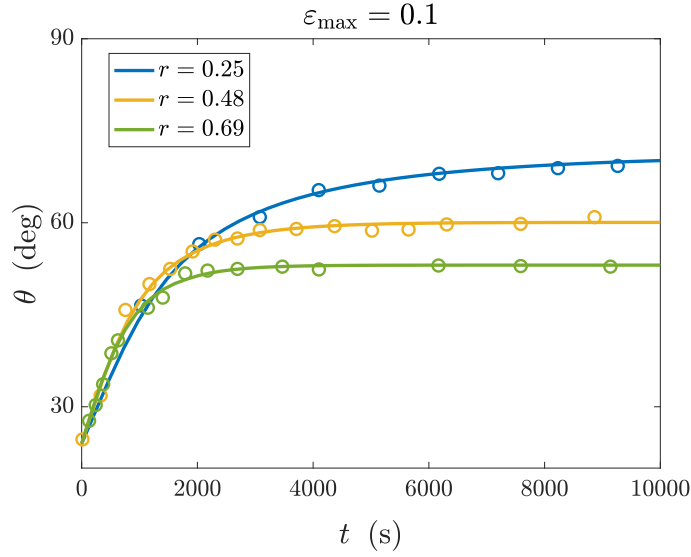


Figure 3: Evolution of the cell orientation angle θ_p with maximum strain of 10%, following Eq. (6.65), for different values of the biaxiality ratio r . The curves show the best fitting of the model, while the dots represent experimental data taken from [22].

- 1 case of equibiaxial deformation when no reorientation occurs. For oblique
- 2 orientations, instead, we have

$$4\kappa^* = 2(1 - r^2)\varepsilon_1^2 \frac{k_4 k_{16} + k_4 k_6 + k_6 k_{14}}{k_4 + k_6} \sin 2\theta_p^*, \quad (6.68)$$

- 3 which is in general different from zero provided that $r \neq 1$.

4 We conclude this Section by comparing the prediction of the constrained
5 model with the data on cell reorientation in [22]. It is seen from Eq. (6.65),
6 that the three parameters \widehat{k}_p , k_m and τ , where the latter is the characteris-

r	k_m/\widehat{k}_p	τ [s]
0.25	0.1594	7.3
0.48	0.2142	5.9
0.69	0.2596	5.7

Table 1: Constitutive parameters in Eq. (6.65) estimated from the fitting of experiments in [22].

1 tic time in \widehat{m} , have to be calibrated. The results of this procedure, carried
2 out through a nonlinear least square algorithm, are shown in Figure 3. The
3 model matches accurately the evolution of the orientation angles seen in the
4 experiments with a relative error below 2 %. The optimal parameters used
5 for the fitting for $r \in \{0.25, 0.48, 0.69\}$ are listed in Table 1 and are indeed
6 coherent with experiments in [22], where the authors found a value of the
7 ratio $k_m/\widehat{k}_p = 0.13 \pm 0.04$ and a characteristic time $\tau = 6.6 \pm 0.4$ s.
8 It is worth noting that the evolution of the angle in Eq. (6.65) only depends
9 on three constitutive parameters including a characteristic time. The eval-
10 uation of the other model coefficients, necessary to estimate the Lagrange
11 multiplier κ^* , would require further experimental data.

12 *Remark.* In the general case of two plane fiber families constrained to remain
13 orthogonal, namely such that $\mathbf{A} + \mathbf{B} = \check{\mathbf{I}}$ with $\check{\mathbf{I}}$ the identity tensor in the
14 fiber plane, it can be easily shown that the stationarity requirement

$$\left[\frac{\partial\phi}{\partial\mathbf{A}}, \mathbf{A}\right]^* + \left[\frac{\partial\phi}{\partial\mathbf{B}}, \mathbf{B}\right]^* = \mathbf{0} \quad (6.69)$$

15 implies

$$[\mathbf{E}, (\phi_4^* - \phi_6^*)\mathbf{A}^* + (\phi_5^* - \phi_7^*)(\mathbf{A}^*\mathbf{E} + \mathbf{E}\mathbf{A}^*)] = \mathbf{0}, \quad (6.70)$$

16 which has the following consequences:

- 17 1. If $\phi_6 = \phi_7 = 0$ identically, meaning that there is a single fiber family,
18 the relation $[\mathbf{E}, \mathbf{S}_c^*] = \mathbf{0}$ for a transversely isotropic material found in
19 [36] is recovered.
- 20 2. If the remodeled direction \mathbf{a} is along a principal direction of strain
21 (and then so is \mathbf{b} because they are orthogonal), the commutator on the
22 l.h.s. of (6.70) is null. Therefore, rotations that align the preferential
23 orientations with the principal strain directions are always stationary
24 solutions of the problem at hand.
- 25 3. If \mathbf{a} is not aligned with a principal strain direction, there might be
26 additional non-trivial solutions identified by the two conditions

$$\phi_4^*(\mathbf{E}, \mathbf{A}^*) = \phi_6^*(\mathbf{E}, \mathbf{A}^*) \quad \text{and} \quad \phi_5^*(\mathbf{E}, \mathbf{A}^*) = \phi_7^*(\mathbf{E}, \mathbf{A}^*).$$

27 6.2. The case of independent reorientation

28 The complex interaction between the cell nucleus and the cytoplasm can
29 be still captured by the proposed model in terms of changes in their relative
30 orientation when cells are cyclically stretched.

Table 2: Constitutive parameters in Eqs. (4.40)-(4.41) identified from the experiments in [29].

k_{14}/k_4	k_{16}/k_6	k_6/k_4	k_8/k_4	τ_p [s]	τ_s [s]
26.5	3.3	9.75	75	128	185

1 In [29], the mechanoadaptive organization of stress fibers and nuclei in ep-
 2 ithelial cells under cyclic stretches is considered, highlighting that these two
 3 cellular components follow different orientation dynamics. In particular, ep-
 4 ithelial cells on a plane substrate were stretched for 2 hours at 5%, 10% and
 5 15% maximum strains whilst measuring the reorientation of the different
 6 stress fiber subtypes. It was seen that dorsal stress fibers, transverse arcs,
 7 and peripheral stress fibers were mainly involved in the cytoplasm response
 8 whereas perinuclear cap fibers were associated with the reorientation and
 9 elongation of the nucleus (see Fig. 2).

10 A sketch of the experimental setup of [29] is shown in Fig. 4(a) where
 11 the coloured arrows are used to indicate the primary (red) and secondary
 12 (green) cell structures, i.e., cytoplasm and nucleus of the cell, respectively.
 13 In the experiments, the longitudinal strain ε_1 was controlled for 2 hours
 14 with the time history shown in inset 4(b) at two different level of maximum
 15 strain (10% and 15%); the lateral strain was also controlled in a way that
 16 $r = -\varepsilon_2/\varepsilon_1 = 0.49$ and the frequency of the deformation was set to 0.3 Hz.
 17 The corresponding reorientation velocities are shown in Fig. 4(c) for the
 18 primary $\dot{\theta}_p$ and the secondary angle $\dot{\theta}_s$. These graphs were obtained by
 19 fitting the experimental data against Eqs. (4.40)-(4.41) (with $m_p = k_4\tau_p$
 20 and $m_s = k_6\tau_s$), through a nonlinear least square algorithm implemented in
 21 MATLAB[®]. The corresponding best fit parameters are listed in Table 2.

22 It is pointed out that the experimental results at 5% were not used for
 23 the fitting process, since they did not show any significant reorientation,
 24 probably caused by the poor interaction between cells and substrate at such
 25 a low level of strain. Finally, Figs. 4(d)-(e) show the evolution of the average
 26 orientation angles during the 2 hours test. [The stretch dependence for both](#)
 27 [the primary and secondary angles is well captured by the model with a](#)
 28 [maximum relative error of 2.5% only seen for the nucleus in inset \(e\).](#) The
 29 fitting is achieved with a value of the constitutive parameter k_8 significantly
 30 larger than k_4 and k_6 , indicating a strong interaction between the two fibre
 31 families.

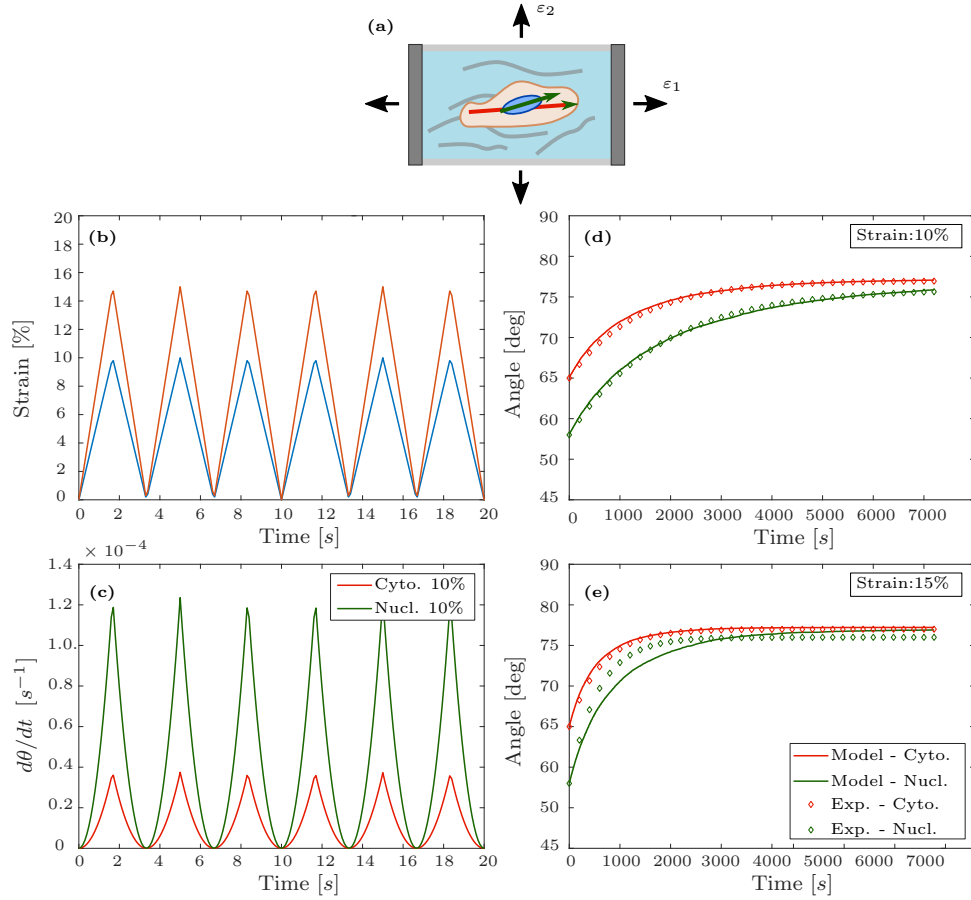


Figure 4: (a): Sketch of the experiments in [29], where the substrate is deformed with a periodic strain. (b): Time history of the longitudinal strain ε_1 imposed to the substrate, with maximum amplitude of 10% (blue curve) and 15% (orange curve). (c): Angular velocity of reorientation for the primary and secondary structures obtained from the model with the optimal estimated constitutive parameters in Tab. 2. (d)–(e): Evolution of the average (over a cycle) orientation angle during the test for the maximum strain 10% (d) and 15% (e), for both the cytoplasm and the nucleus, compared with experimental results in [29].

1 7. Conclusions

2 We presented a model to describe cell alignment under applied trac-
3 tion using a fiber reorientation framework, in which two different families
4 of fibers are able to rotate independently under the action of two distinct
5 rotation tensors. The framework accounts for the elastic coupling between
6 the fiber families, while the introduction of a proper constraint allowed us
7 to recover some literature results on reorientation of transversely isotropic
8 and orthotropic materials. The stationary solutions of the remodelling equa-
9 tions are then thoroughly studied to find a generalization of a well-known
10 coaxiality theorem by Vianello, that holds in the passive case; active effects,
11 even if not studied in details in the present paper, can be naturally and
12 straightforwardly incorporated in the modelling equations.

13 As a main application of the proposed framework, we studied cell reorienta-
14 tion under stretch. Perfect adhesion is assumed between the cell layer and
15 the substrate and the rotations of the cells due to mechanical stimuli are
16 viewed as additional state variables of the problem. By using experimental
17 data available from the literature, we have found that the model is able to
18 accurately match the experimental results on cell orientation for both the
19 constrained and unconstrained fiber evolutions. Even though the focus is
20 mainly on biological applications, the proposed framework is indeed general.
21 As such, it could be readily adapted to describe engineering materials like
22 composites with two families of fibers [54], which are engineered to reorient
23 under different stimuli.

24 The thermodynamically consistent structure of the model offers advantages
25 over the existing phenomenological approaches pursued in the literature and
26 may open the road to future developments. These include: the possibility
27 of having a mobility tensor dependent both on time and on strain so to
28 describe the dynamic response seen for certain types of cells [55, 56]; the
29 incorporation of a non-perfect adhesion between cells and substrate by in-
30 ducing reorientation only when a certain amount of strain is accumulated
31 [55] or when adhesion bonds are broken as a consequence of mechanical de-
32 formation; the inclusion of the matrix viscoelasticity, that in general induces
33 additional characteristic times in the material response. The hierarchic role
34 of the fiber families may also be incorporated by distinguishing between
35 “pulled” and “dragged” reorientations, that occur when one fiber family
36 drives the reorientation of the other through the elastic coupling.

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22 **Appendix A. An alternative choice for the constraint equation**

23 Here we consider the alternative choice of the scalar constraint

$$c(\mathbf{A}, \mathbf{B}) = \mathbf{A} \cdot \mathbf{B} - \mathbf{A}_0 \cdot \mathbf{B}_0, \quad (\text{A.1})$$

24 which gives

$$\dot{c}(\mathbf{A}, \mathbf{B}) = \dot{\mathbf{A}} \cdot \mathbf{B} + \mathbf{A} \cdot \dot{\mathbf{B}} = [\mathbf{A}, \mathbf{B}] \cdot (\dot{\mathbf{R}}_s \mathbf{R}_s^T - \dot{\mathbf{R}}_p \mathbf{R}_p^T). \quad (\text{A.2})$$

25 Accordingly, following the procedure detailed in Section 5, the evolution
26 equations of the constrained remodeling problem are written as

$$\left\{ \begin{array}{l} \mathbb{D}_p \dot{\mathbf{R}}_p \mathbf{R}_p^T = -\left[\frac{\partial \phi}{\partial \mathbf{A}}, \mathbf{A}\right] + \kappa [\mathbf{A}, \mathbf{B}], \\ \mathbb{D}_s \dot{\mathbf{R}}_s \mathbf{R}_s^T = -\left[\frac{\partial \phi}{\partial \mathbf{B}}, \mathbf{B}\right] - \kappa [\mathbf{A}, \mathbf{B}], \\ \mathbf{A} \cdot \mathbf{B} = \mathbf{A}_0 \cdot \mathbf{B}_0, \end{array} \right. \quad (\text{A.3})$$

- 1 having introduced a scalar Lagrange multiplier κ . At stationarity, summing
 2 the two equations one gets

$$\left[\frac{\partial\phi}{\partial\mathbf{A}}, \mathbf{A}\right]^* + \left[\frac{\partial\phi}{\partial\mathbf{B}}, \mathbf{B}\right]^* = \mathbf{0} = [\mathbf{E}, \mathbf{S}_c]^* \quad (\text{A.4})$$

- 3 and subtracting instead yields

$$\kappa^* [\mathbf{A}, \mathbf{B}]^* = \left[\frac{\partial\phi}{\partial\mathbf{A}}, \mathbf{A}\right]^*. \quad (\text{A.5})$$

- 4 On assuming that $\mathbf{a} = \cos\theta_p\mathbf{e}_1 + \sin\theta_p\mathbf{e}_2$ and $\mathbf{b} = \cos\theta_s\mathbf{e}_1 + \sin\theta_s\mathbf{e}_2$, the
 5 constraint equation (A.1) gives

$$c(\mathbf{a}, \mathbf{b}) = \cos^2(\theta_p - \theta_s) - \cos^2(\theta_{p0} - \theta_{s0}), \quad (\text{A.6})$$

whence the remodeling equations become

$$\begin{cases} 2m_p\dot{\theta}_p = (1+r)\varepsilon_1^2 F(\theta_p, r; k_4, k_{14}) + \frac{1}{2}\varepsilon_1^2 k_8 [G(\theta_p, \theta_s, r) + H(\theta_p, \theta_s, r)] + \kappa \sin(2(\theta_s - \theta_p)) \\ 2m_s\dot{\theta}_s = (1+r)\varepsilon_1^2 F(\theta_s, r; k_6, k_{16}) + \frac{1}{2}\varepsilon_1^2 k_8 [G(\theta_s, \theta_p, r) + H(\theta_p, \theta_s, r)] - \kappa \sin(2(\theta_s - \theta_p)) \\ \cos^2(\theta_p - \theta_s) = \cos^2(\theta_{p0} - \theta_{s0}). \end{cases} \quad (\text{A.7})$$

For an equibiaxial test, with $r = -1$, by summing and subtracting the first two equations in (A.7) one obtains

$$\begin{cases} m_p\dot{\theta}_p + m_s\dot{\theta}_s = 0 \\ m_p\dot{\theta}_p - m_s\dot{\theta}_s = \varepsilon_1^2 k_8 \cos^2(\theta_p - \theta_s) \sin(2(\theta_p - \theta_s)) - \kappa \sin(\theta_p - \theta_s) \\ \cos^2(\theta_p - \theta_s) = \cos^2(\theta_{p0} - \theta_{s0}). \end{cases} \quad (\text{A.8})$$

- 6 The constraint equation $\cos^2(\theta_p - \theta_s) = \pm \cos^2(\theta_{p0} - \theta_{s0})$ implies that

$$\dot{c} = \sin(2(\theta_s - \theta_p))(\dot{\theta}_p - \dot{\theta}_s) = 0, \quad (\text{A.9})$$

- 7 which has the solutions $\dot{\theta}_p = \dot{\theta}_s$ or $\sin(2(\theta_s - \theta_p)) = 0$, that is $\Delta\theta = \theta_s -$
 8 $\theta_p = (1+k)\frac{\pi}{2}$, $\kappa = 0, 1, 2, \dots$. The former substituted into (A.8) gives
 9 $\dot{\theta}_p = \dot{\theta}_s = 0$ and $\kappa = \varepsilon_1^2 k_8 \cos^2(\Delta\theta_0)$. The latter gives $\dot{\theta}_p = \dot{\theta}_s = 0$ and an
 10 indeterminate reaction couple κ . In both cases, however, it is obtained that
 11 no remodeling occurs when equibiaxial strains are considered, as already
 12 seen from Eq. (5.59).