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## NON-LINEAR MODEL FOR COMPRESSION TESTS **ON ARTICULAR CARTILAGE**

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ABSTRACT

Hydrated soft tissues, such as articular cartilage, are often modelled as biphasic systems with individually incompressible solid and fluid phases, and biphasic models are employed to fit experimental data in order to determine the mechanical and hydraulic properties of the tissues. Two of the most common experimental setups are confined and unconfined compression. Analytical solutions exist for the unconfined case with the linear, isotropic, homogeneous model of articular cartilage, and for the confined case with the non-linear, isotropic, homogeneous model. The aim of this contribution is to provide an easily implementable numerical tool to determine a solution to the governing differential equations of (homogeneous and isotropic) unconfined and (inhomogeneous and isotropic) confined compression under large deformations. The large-deformation governing equations are reduced to equivalent diffusive equations, which are then solved by means of Finite Difference methods. The solution strategy proposed here could be used to generate benchmark tests for validating complex user-defined material models within Finite Element implementations, and for determining the tissue's mechanical and hydraulic properties from experimental data.

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## 1 Introduction

Since its introduction, the biphasic model of articular car-2 tilage [1–3] has been the standard manner to study also other hydrated soft tissues. In this model, cartilage is represented as the mixture of an incompressible solid, representing structural 5 macromolecules such as collagen fibres and proteoglycans, and an incompressible fluid, representing the interstitial water, along with the various chemical species dissolved in it. In order to fully characterise the behaviour of cartilage according to the biphasic model, it is necessary to experimentally evaluate its elastic properties and its permeability (which is the parameter accounting for the solid-fluid interaction). The most common tests are confined and unconfined compression. In the former, a cartilage sample is placed in an impermeable, rigid chamber and compressed by a porous, rigid piston, so that the fluid can escape from the sample through the piston. In the latter, cartilage is squeezed between two impermeable, rigid plates, so that it can freely expand laterally and fluid can freely escape through the lateral boundary.

Aside from many studies based on Finite Element Analysis, 19 confined and unconfined compression have been also modelled 20 analytically in some particular cases. Based on the linear bipha-21 sic model of articular cartilage [1-3], Armstrong et al. [4] derived 22

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an analytical solution for the unconfined compression test under 23 small deformations, in terms of series expansions. Holmes and 24 Mow [5] proposed an isotropic homogeneous model of articular 25 cartilage, with non-linear elasticity and deformation-dependent 26 permeability, and studied the case of confined compression ana-27 lytically. Moreover, in a previous work [6], unconfined compres-28 sion has been solved numerically under small deformations for 29 a linear, isotropic, inhomogeneous model. However, these spe-30 cific cases cannot be used to describe many experimental set-up 31 conditions. 32

In this work, based on the large-strain governing equations 33 of a biphasic mixture (e.g., [7–9]), we propose a solution to 34 the differential equations of both unconfined and confined com-35 pression problems under large deformations, with isotropic nonlinear elasticity and deformation-dependent permeability [5]. 37 The case of unconfined compression is studied under the hypoth-38 esis of homogeneity, whereas in that of confined compression the 39 elastic properties and permeability are inhomogeneous, as ob-40 tained from published experimental works [10, 11], and similarly 41 to what has been done in [6] for the small-deformation case. 42

Once the hyperelastic constitutive equations are set, the gov-43 erning equations consist of a system of 4 differential equations in 44 4 unknowns: three components of the configuration map (treated 45 in terms of their derivatives, i.e., the components of the defor-46 mation gradient tensor), and the fluid pressure. For the cases of 47 homogeneous unconfined compression and inhomogeneous con-48 fined compression, the material gradient of the pressure is elimi-49 nated, yielding a single scalar equation in the volume ratio of the 50 diffusion-advection type, which simplifies remarkably the math-51 ematical problem. The solution that we propose is obtained nu-52 merically via a direct application of Finite Difference schemes, 53 and can be used as a rapid, yet effective, comparison solution 54 to verify the robustness and accuracy of complex user-defined material models within Finite Element methods. Furthermore, 56 the proposed implementation is easily manageable and makes 57 the code potentially useful in the determination of mechanical 58 parameters, directly fitting experimental curves. 59

#### 2 **Balance Laws** 60

Here, the description of articular cartilage is limited to the 61 macro-scale, which is interpreted as the laboratory scale at which 62 constitutive information on the overall mechanical behaviour of 63 a given sample of tissue is extracted by means of experiments. At 64 this scale, the tissue is modelled as a biphasic mixture comprising 65 a solid and a fluid phase. The solid phase is the macro-scale rep-66 resentation of a deformable porous medium, which, in fact, is it-67 self a mixture composed mainly of proteoglycans and collagen fi-68 bres. The fluid phase represents the interstitial fluid, which occu-69 pies the voids of the porous medium and consists mainly of wa-70 ter, ions and various types of chemical compounds, such as nutri-71 ents for the cells and byproducts of the cellular metabolism [12]. 72

In the following, the subscripts 's' and 'f' shall specify 73 74 whether a given physical quantity is associated with the solid or with the fluid phase. When there is no danger of confusion, the terms "phase" and "constituent" shall be used interchangeably. The mass distribution of the  $\alpha$ th phase of the mixture ( $\alpha = f, s$ ), can be expressed either per unit volume occupied by the  $\alpha$ th phase itself, or per unit volume of the mixture as a whole. In the first case, one speaks of the "true", or intrinsic, mass density 80  $\rho_{\alpha}$  of the  $\alpha$ th phase. In the second case, instead, one introduces 81 the "apparent" mass density  $\phi_{\alpha}\rho_{\alpha}$ , with  $\phi_{\alpha}$  being the volumetric fraction of the  $\alpha$ th phase, i.e., the ratio between the size of 84 the volume occupied by  $\alpha$ th phase and the size of a representa-85 tive volume for the mixture as a whole. Note that the mixture is subjected to the saturation constraint  $\phi_s + \phi_f = 1$ . All the balance laws referred to the macro-scale description of the mixture's constituents are formulated by employing the apparent mass densities of the phases.

In the absence of sources and sinks of mass, the spatial, local form of the mass balance laws associated with the fluid and the solid phase can be written as

$$\partial_t(\phi_f \rho_f) + \operatorname{div}(\phi_f \rho_f \boldsymbol{v}_s) + \operatorname{div}(\rho_f \boldsymbol{w}) = 0, \qquad (1a)$$

$$\partial_t(\phi_{\rm s}\rho_{\rm s}) + {\rm div}(\phi_{\rm s}\rho_{\rm s}\boldsymbol{\nu}_{\rm s}) = 0, \qquad (1b)$$

where  $v_s$  and  $v_f$  are the velocities of the solid and fluid phase, 93 respectively, and  $\mathbf{w} = \phi_f(\mathbf{v}_f - \mathbf{v}_s)$  is the filtration velocity. For 94 more details on the kinematics of biphasic mixtures, see, for in-95 96 stance, [13]. From here on, the mass densities  $\rho_{\rm f}$  and  $\rho_{\rm s}$  are assumed to be given constants, which means that both the fluid and 97 the solid phases are regarded as intrinsically incompressible ma-98 terials. By definition, this means that the substantial derivatives 99  $D_{\alpha}\rho_{\alpha} := \partial_t \rho_{\alpha} + (\operatorname{grad} \rho_{\alpha}) \boldsymbol{v}_{\alpha}$  are zero for  $\alpha = f, s$ . 100

Under the assumption of negligible inertial effects, in the absence of body forces external to the considered mixture, and accepting the validity of Darcy's law, the spatial, local balance laws of momentum associated with the mixture as a whole and the fluid phase can be written as

$$\mathbf{0} = \operatorname{div}(\boldsymbol{\sigma}_{\mathrm{f}} + \boldsymbol{\sigma}_{\mathrm{s}}), \qquad (2)$$

$$\boldsymbol{w} = -\boldsymbol{k} \operatorname{grad} \boldsymbol{p}, \qquad (3)$$

where  $\boldsymbol{\sigma}_{f}$  and  $\boldsymbol{\sigma}_{s}$  are the Cauchy stress tensors of the fluid and solid phase, respectively, and  $\boldsymbol{k}$  is the spatial permeability tensor. The mixture is assumed to be closed with respect to momentum. 108

If the fluid phase is modelled as an incompressible and macroscopically inviscid Stokes fluid, the stress tensors for the

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fluid, the solid and the whole tissue admit the expressions

$$\sigma_{\rm f} = -\phi_{\rm f} p g^{-1},$$
 (4a) <sup>146</sup>

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$$\boldsymbol{\sigma}_{\rm s} = -\phi_{\rm s} \, p \, \boldsymbol{g}^{-1} + \boldsymbol{\sigma}_{\rm c} \,, \tag{4b} \quad {}_{148}$$

$$\boldsymbol{\sigma} = \boldsymbol{\sigma}_{\rm f} + \boldsymbol{\sigma}_{\rm s} = -p \boldsymbol{g}^{-1} + \boldsymbol{\sigma}_{\rm c}, \qquad (4c)$$

where  $\boldsymbol{\sigma}_{c}$  is referred to as the *constitutive part* of  $\boldsymbol{\sigma}_{s}$ , *p* is the hy-112 drostatic pressure, and  $g^{-1}$ , with components  $g^{ab}$ , is the inverse 113 of the spatial metric tensor g and serves here as the "contravari-114 ant" identity tensor. 115

The deformation of the solid phase is denoted by  $\chi$ , the de-116 formation gradient by **F** (with components  $F^a{}_A = \chi^a{}_A$ ), and the 117 volume ratio by  $J = \det F$ . By performing a backward Piola 118 Transformation of (1a) and (1b), which is done by multiplying 119 both equations by J, one obtains 120

$$\dot{\phi}_{\mathrm{fR}} + \mathrm{Div}(\boldsymbol{W}) = 0, \qquad (5a)$$

$$\dot{\phi}_{\rm sP} = 0,$$
 (5a) (5a) (5a) (5b) 157

In (5a) and (5b), the superimposed dot stands for time differenti-121 ation, Div is the material divergence operator, and 122

$$\phi_{\rm sR} = J\phi_{\rm s}\,,\tag{6a}$$

$$\phi_{\rm fR} = J\phi_{\rm f} = J - \phi_{\rm sR} \,, \tag{6b}$$

$$\boldsymbol{W} = J\boldsymbol{F}^{-1}\boldsymbol{w} = -\boldsymbol{K}\operatorname{Grad} p, \qquad (6c) \quad {}_{161}$$

with  $\mathbf{K} = J\mathbf{F}^{-1}\mathbf{k}\mathbf{F}^{-T}$  being the material permeability tensor, are 123 the Piola transforms of  $\phi_f$ ,  $\phi_s$  and w. In particular, (6a) can 124 be used to express  $\phi_s$  as a function of the volume ratio of the 125 solid phase, i.e.,  $\phi_{\rm s} = J^{-1}\phi_{\rm sR}$ . This result, which stems from 126 the incompressibility of the solid phase, permits to rephrase the 127 inequalities  $0 \le \phi_s(x,t) \le 1$  (with the upper bound condition 128  $\phi_{\rm s}(x,t) = 1$  implying that the limit of *compaction* is reached) 129 as  $0 \le \phi_{sR}(X) \le J(X,t)$ , and thus it places on J the unilateral 130 constraint  $J(X,t) \ge \phi_{\rm sR}(X)$  [8]. In particular, when the condition 131  $J = \phi_{sR}$  is met at a given point *X* of the reference configuration 132  $\mathcal{B}_{R}$ , all fluid has been expelled from the point, which thereby re-133 mains composed of solid alone, which is incompressible by hy-134 pothesis. It is worth to recall that, for a biphasic mixture, the re-135 quirement that both phases are intrinsically incompressible, does 136 *not* lead to the restriction J = 1 of isochoric motion, due to the 137 presence of the volumetric fraction  $\phi_s$  in (1b). Indeed, the as-138 sumption of incompressibility, which is translated into  $D_{\rm s}\rho_{\rm s}=0$ , 139 transforms (1b) into an equation for  $\phi_s$ , whose variations are 140 compensated for by the change of volume of the solid phase. In 141 the material formalism, this fact is reflected by (6a), which allows 142 to express  $\phi_s$  as a function of J. An extensive discussion about 143

this issue and, in particular, on the consequences of compaction, is given in [8].

Finally, by adding together (5a) and (5b), using (6c), and performing a Piola transformation of (2), with  $\sigma_{\rm f}$  and  $\sigma_{\rm s}$  given by (4b) and (4a), the material form of the mass and momentum balance laws becomes

$$\dot{J} = \operatorname{Div}\left(\boldsymbol{K}\operatorname{Grad} p\right), \qquad (7a)$$

$$\operatorname{Div} \boldsymbol{P}_{c} = J \boldsymbol{g}^{-1} \boldsymbol{F}^{-T} \operatorname{Grad} \boldsymbol{p}, \qquad (7b)$$

where  $P_{c} = J \sigma_{c} F^{-T}$  is the constitutive part of the first Piola-Kirchhoff stress tensor of the solid phase. Furthermore, the first Piola-Kirchhoff stress for the whole tissue is obtained by Piolatransforming (4c), which yields  $\boldsymbol{P} = -J p \boldsymbol{g}^{-1} \boldsymbol{F}^{-T} + \boldsymbol{P}_{c}$ .

#### Constitutive Laws and Final Model Equations 3

The non-linear isotropic model proposed by Holmes and Mow [5] is adopted in this work. The solid phase is regarded as hyperelastic, with potential

$$\hat{W}(\boldsymbol{C}) = \boldsymbol{\alpha}_0 \left( \exp[\boldsymbol{\varphi}(\boldsymbol{C})] - 1 \right), \tag{8a}$$

$$\varphi(\boldsymbol{C}) = \alpha_1 \left[ I_1(\boldsymbol{C}) - 3 \right] + \alpha_2 \left[ I_2(\boldsymbol{C}) - 3 \right] - \beta \ln \left[ I_3(\boldsymbol{C}) \right], \quad (8b)$$

where  $\alpha_0, \alpha_1, \alpha_2$ , and  $\beta$  are material parameters, and  $I_1(\mathbf{C}) =$  $tr(C), I_2(C) = \frac{1}{2}[(tr(C))^2 - tr(C^2)], and I_3(C) = det(C) are the in$ variants of the right Cauchy-Green deformation tensor  $\boldsymbol{C} = \boldsymbol{F}^{\mathrm{T}} \boldsymbol{F}$ . Thus,  $P_c$  is given by

$$\boldsymbol{P}_{c} = \hat{\boldsymbol{P}}_{c}(\boldsymbol{F}) = \boldsymbol{F}\left(2\frac{\partial\hat{W}}{\partial\boldsymbol{C}}(\boldsymbol{C})\right).$$
(9)

The permeability is assumed to be related to J via the Holmes-162 Mow law [5] 163

$$k = \hat{k}(J) = k_0 \left(\frac{J - \phi_{sR}}{1 - \phi_{sR}}\right)^{\gamma} \exp\left(\frac{M}{2}(J^2 - 1)\right), \quad (10)$$

where  $\gamma$  and M are material parameters,  $\hat{k}(J) = k$  denotes the 165 constitutive function associated with the scalar permeability k, and  $k_0 = \hat{k}(1)$  is the value of the permeability in the undeformed 166 configuration (J = 1). In order to satisfy (7a),  $\hat{k}$  must vanish 167 at compaction, i.e., at  $J = \phi_{sR}$ , so that  $\dot{J}$  vanishes too, and the incompressibility constraint is respected. As an isotropic tensorvalued function, the permeability is assumed to be spherical [14], so that the spatial and material permeability tensors are given by

$$\mathbf{k} = k\mathbf{g}^{-1} = \hat{k}(J)\mathbf{g}^{-1},$$
 (11a)

$$\boldsymbol{K} = \hat{\boldsymbol{K}}(\boldsymbol{C}) = J\,\hat{k}(J)\,\boldsymbol{C}^{-1}.$$
(11b)

In (10),  $\hat{k}$  vanishes for  $J = \phi_{sR}$ , and therefore **k** vanishes too. 172 For an inhomogeneous material,  $\hat{W}$  and  $\hat{K}$  depend explicitly on 173 the material point X, through the parameters  $\alpha_0, \alpha_1, \alpha_2, \beta$  and 174  $k_0, \gamma, M$ , and possibly through  $\phi_{sR}$  as well. Hereafter, cartilage 175 is regarded as homogeneous for the case of unconfined compres-176 sion and as inhomogeneous for the case of confined compression. 177 Equations (7a) and (7b) are suitable for computations based 178 on the Finite Element Method (FEM), cf., e.g., [12]. Here, how-179 ever, a different approach is followed, since the aim of this work 180

is to provide a valid alternative to Finite Element implementa-181 tions for the considered problems. The reason for undertaking 182 this task is to supply fast estimates about the hydraulic and me-183 chanical properties of cartilage (in the limit case of isotropy), 184 that can be used as reference for testing the reliability of com-185 plex, FEM-based numerical strategies, which might be necessary 186 for highly non-linear, coupled, anisotropic and inhomogeneous 187 problems. The first step is the decoupling of (7a) from (7b), 188 which is achieved by substituting  $\operatorname{Grad} p$ , obtained from (7b), 189 into (7a). By accounting for (11b), this yields 190

$$\vec{J} = \operatorname{Div}\left[\hat{k}(J)\boldsymbol{F}^{-1}\operatorname{Div}\left(\boldsymbol{\hat{P}}_{c}(\boldsymbol{F})\right)\right], \qquad (12a)$$

Grad 
$$p = J^{-1} \boldsymbol{F}^{\mathrm{T}} \boldsymbol{g} \operatorname{Div}(\hat{\boldsymbol{P}}_{\mathrm{c}}(\boldsymbol{F}))$$
. (12b)

Equations (12a) and (12b) have some relevant differences 191 with respect to the original Equations (7a) and (7b). Firstly, p can 192 be computed *a posteriori* by solving (12b), once (12a) is solved 193 for  $\chi$ . Secondly, (12b) involves only the first-order space deriva-194 tives of p, whereas the Poisson-like equation (7a) also involves 195 the second-order space derivatives of p. Finally, the permeability 196 does not directly affect the pressure. Rather, it influences the so-197 lution of (12a), which involves the third-order space derivatives 198 of  $\chi$  as well as its mixed derivatives (i.e., with respect to both 199 time and space), as prescribed by the computation of J. 200

### 201 4 Axisymmetric Unconfined Compression

The subject of this section is the study of the unconfined compression test of a cylindrical specimen of cartilage. In this test, the specimen is assumed to be homogeneous and isotropic. Therefore, its permeability and hyperelastic potential are independent of material points, and related to deformation only through the invariants of C. Moreover,  $\phi_{sR}$  is a model constant.

The cylindrical specimen is inserted between two rigid and 208 impermeable plates that remain parallel to each other for the 209 whole duration of the experiment. The lower plate is kept fixed, 210 while the upper one moves downward according to a prescribed 211 loading protocol. In this work, only a displacement-control test is 212 considered. The lateral wall of the specimen is traction-free and 213 permeable. The lower and upper surfaces of the specimen are 214 allowed to glide on the lower and upper plate, respectively, in an 215 axisymmetric way. Moreover, no friction is considered, so that 216

the specimen preserves its original cylindrical shape throughout the experiment.

The geometry of the specimen, its material symmetries (homogeneity and isotropy) and the experimental protocol make it convenient to employ cylindrical coordinates  $\{R, \Theta, Z\}$  and  $\{r, \theta, z\}$  for both the reference (undeformed) and deformed configuration, respectively. Below, the boundary conditions for  $\chi$ and p, which have to hold at all times, are specified for all portions of the boundary.

At the lower boundary,  $(R, \Theta, Z) \in [0, R_{\text{ext}}] \times [0, 2\pi[\times \{0\},$ 

$$\chi^z = 0,$$
 [no displacement] (13a)  
(-**K**Grad p).(-**E**<sub>Z</sub>) = 0, [no flux] (13b)

where  $\chi^z$  is the axial component of the deformation  $\chi$ , and  $E_Z$  is the unit vector pointing upward and aligned along the axial direction.

At the upper boundary,  $(R, \Theta, Z) \in [0, R_{\text{ext}}] \times [0, 2\pi[\times \{H\},$ 

$$\chi^z = \lambda_Z H \,, \tag{14a}$$

$$\lambda_Z(t) = 1 - \frac{u_T}{H} \left[ 1 - \exp(-t/t_u) \right], \quad \text{[prescribed stretch]} \quad (14b)$$

$$(-\boldsymbol{K}\operatorname{Grad} p) \cdot \boldsymbol{E}_Z = 0,$$
 [no flux] (14c)

where *H* is the initial height of the specimen, and  $\lambda_Z$  is the imposed time-dependent stretch, with target displacement  $u_T$  and time constant  $t_u$ .

At the lateral boundary,  $(R, \Theta, Z) \in \{R_{ext}\} \times [0, 2\pi[\times[0, H]],$ 

$$-p = 0$$
, [atmospheric pressure] (15a)

$$\boldsymbol{P}.\boldsymbol{E}_{R} = \boldsymbol{0},$$
 [traction-free boundary] (15b)

where  $E_R$  is the referential radial unit vector, pointing outward and aligned along the radial direction. Finally, the axial symmetry of the problem places the further restriction that the radial deformation and the radial fluid flux must vanish at the origin of each cross section of the specimen. Since the reference configuration is assumed to coincide with the stress-free, undeformed one, the initial conditions p(X,0) = 0 and  $\chi(X,0) = X$  apply at all inner points X of the computational domain.

It should be remarked that (15b) involves the overall first Piola-Kirchhoff stress tensor of the mixture as a whole. Since it holds that  $\mathbf{P} = -J p \mathbf{g}^{-1} \mathbf{F}^{-T} + \mathbf{P}_c$ , and pressure has to vanish on the lateral boundary of the specimen, (15b) can also be rephrased in terms of the constitutive part of  $\mathbf{P}$ , i.e.  $\mathbf{P}_c \cdot \mathbf{E}_R = \mathbf{0}$ .

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#### 248 4.1 Specific Form of the Deformation

<sup>249</sup> Due to the symmetries of the problem,  $\chi$  acquires the form

$$r = \chi^{r}(R, \Theta, Z, t) \equiv f(R, t),$$
 (16a)

$$\vartheta = \chi^{\vartheta}(R, \Theta, Z, t) = \Theta, \qquad (16b)$$

$$z = \chi^{z}(R, \Theta, Z, t) = \lambda_{Z}(t) Z.$$
 (16c)

In (16a),  $\chi^r$  is re-defined as a function f of R and t alone, and <sup>251</sup>  $\lambda_Z(t)$  is the uniform axial stretch, as defined in (14b). The latter <sup>252</sup> is a function known from the boundary conditions on the dis-<sup>253</sup> placement in the axial direction for the case of a displacement-<sup>254</sup> controlled test. The stretches in the radial and circumferential <sup>255</sup> direction are given by

$$\lambda_{R}(R,t) = \frac{\partial f}{\partial R}(R,t) \equiv f'(R,t), \qquad (17a)$$

$$\lambda_{\Theta}(R,t) = \frac{f(R,t)}{R}.$$
(17b)

Thus, the matrix representation of F, which is diagonal, and the volume ratio J become

$$[F^{a}_{A}](R,t) = \operatorname{diag}[\lambda_{R}(R,t), \lambda_{\Theta}(R,t), \lambda_{Z}(t)], \qquad (18a)$$

$$J(R,t) = f'(R,t) \frac{f(R,t)}{R} \lambda_Z(t).$$
(18b)

## 258 4.2 Stress and Balance Equations

<sup>259</sup> Because of the deformation specified by (16a)–(16c), the <sup>260</sup> matrix representation of  $P_c$  is diagonal. Moreover, the equation <sup>261</sup> of the balance of mass (12a), and the only non-trivially satisfied <sup>262</sup> component of the equation of balance of momentum (12b) read

$$\dot{J} = \left(\frac{\partial}{\partial R} + \frac{1}{R}\right) \left[\frac{\hat{k}(J)}{\lambda_R} \left(\frac{\partial P_c^{rR}}{\partial R} + \frac{P_c^{rR} - P_c^{\vartheta\Theta}}{R}\right)\right], \quad (19a) \quad {}^{290}_{291}$$

$$\frac{\partial p}{\partial R} = \frac{\lambda_R}{J} \left( \frac{\partial P_c^{rR}}{\partial R} + \frac{P_c^{rR} - P_c^{\vartheta \Theta}}{R} \right).$$
(19b)

Since  $P_c^{rR}$  and  $P_c^{\vartheta\Theta}$  are constitutive functions of  $\lambda_R$ ,  $\lambda_{\Theta}$  and  $\lambda_Z$ , and since  $\lambda_R$  and  $\lambda_{\Theta}$  involve the radial deformation f, while  $\lambda_Z$  is known from the outset, the right-hand-side of (19a) can be recast as a combination of terms in the unknown f and its radial derivatives up the third-order. Thus, after substituting the constitutive laws, an equation for f can be obtained.

Since (19b) is decoupled from (19a), it suffices to determine f by solving (19a) and then compute *p* through (19b).

#### 271 4.3 "Diffusive Equation"

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Solving (19a) may be cumbersome, since it is a highly nonlinear partial differential equation of the third-order in the radial derivatives of f, and it involves the mixed derivatives of fwith respect to time and the radial coordinate. The scope of this section is to show that (19a) can be transformed into a pseudodiffusion-reaction equation in J. To achieve this goal, the first step consists of the change of variables

$$f'(R,t) = \lambda_R(R,t) = \frac{RJ(R,t)}{f(R,t)\lambda_Z(t)}.$$
(20)

Accordingly,  $\lambda_R$  can be viewed as a function of J, f,  $\lambda_Z$  and R, where the dependence on  $\lambda_Z(t)$ , which is known from the outset, can be rephrased as an explicit dependence on time. Similarly,  $\lambda_{\Theta}$  can be regarded as a function of f and R. Hence, the stresses  $P_c^{R}$  and  $P_c^{\partial \Theta}$  can be reformulated as follows:

$$P_{\rm c}^{rR} = \tilde{P}_{\rm c}^{rR}(J(R,t), f(R,t), \lambda_Z(t), R), \qquad (21a)$$

$$P_{\rm c}^{\vartheta\Theta} = \tilde{P}_{\rm c}^{\vartheta\Theta}(J(R,t), f(R,t), \lambda_Z(t), R).$$
(21b)

By substituting the right-hand-sides of (21a) and (21b) into (19a), and performing some algebraic manipulations that account for the new definitions of stress (21a) and (21b), it is possible to define the quantities

$$D := \frac{k}{\lambda_R} \frac{\partial \tilde{P}_c^{rR}}{\partial J}, \qquad (22a)$$

$$-\mathcal{A}J := \frac{k}{\lambda_R} \left\{ \frac{\partial \tilde{P}_c^{rR}}{\partial f} \lambda_R + \frac{\partial \tilde{P}_c^{rR}}{\partial R} \bigg|_{\exp} + \frac{\tilde{P}_c^{rR} - \tilde{P}_c^{\vartheta\Theta}}{R} \right\}, \quad (22b)$$

where  $[\partial(\cdot)/\partial R]\Big|_{exp}$  represents the explicit derivative along the radial direction. Consequently, (19a) takes the form of a diffusion-advection equation in the variable *J* (the transported field), i.e.,

$$\dot{J} = \left(\frac{\partial}{\partial R} + \frac{1}{R}\right) \left[\mathcal{D}\frac{\partial J}{\partial R} - \mathcal{A}J\right],$$
(23)

with  $\mathcal{D}$  and  $\mathcal{A}$  playing the role of the diffusion coefficient and advection velocity, respectively. The physical units of  $\mathcal{D}$  and  $\mathcal{A}$ , which are given by  $[\mathcal{D}] = \text{length}^2/\text{time}$  and  $[\mathcal{A}] = \text{length}/\text{time}$ , show that these identifications are physically sound.

It is worth to mention that  $\mathcal{D}$  stems from the combination of very important physical entities. These are the permeability, which encapsulates all information about the hydraulic response of the system, and the derivative of  $\tilde{P}_c^{rR}$  with respect to J, which

is related to the acoustic tensor of the solid phase. Analogous 300 331 considerations hold true for the drift velocity A. In this case, 301 however, also the term  $(\tilde{P}_{c}^{rR} - \tilde{P}_{c}^{\vartheta\Theta})/R$  contributes to advection. 302 The coefficients  $\mathcal{D}$  and  $\mathcal{A}$  can be expressed as functions of J, 303 f,  $\lambda_Z$  and R. Therefore, the diffusion-advection equation (23) is 304 coupled with the radial deformation f, which can be determined 305 by solving (20). In conclusion, the change of variables (20) 306 rephrases the mathematical structure of (19a) and (19b) into the 307 following set of new model equations: 308

$$f'(R,t) = \frac{RJ(R,t)}{f(R,t)\lambda_Z(t)},$$
(24a)

$$\dot{J} = \frac{1}{R} \frac{\partial}{\partial R} \left\{ R \left[ \mathcal{D} \frac{\partial J}{\partial R} - \mathcal{A} J \right] \right\}, \qquad (24b)$$

$$\frac{J}{\lambda_R}\frac{\partial p}{\partial R} = \frac{\lambda_R}{\hat{k}(J)} \left[ \mathcal{D}\frac{\partial J}{\partial R} - \mathcal{A}J \right].$$
(24c)

This set consists of three independent scalar equations in the 309 three unknowns J, f and p. Clearly, the boundary conditions 310 must be rewritten accordingly: 311

$$f(0,t) = 0$$
 [axial symmetry], (25a)

$$\left(\mathcal{D}\frac{\partial J}{\partial R} - \mathcal{A}J\right)\Big|_{R=0} = 0 \quad \text{[axial symmetry]}, \quad (25b) \quad {}^{341}_{342}$$

$$p(R_{\text{ext}},t) = 0$$
 [from (15a)], (25c)

$$\tilde{P}_{c}^{rR}(J, f, \lambda_{Z}(t), R)\Big|_{R=R_{\text{ext}}} = 0 \qquad [\text{from (15b)}]. \quad (25d)$$

Note that (25a) is a homogeneous Dirichlet condition on f (only 312 one boundary condition is needed for f, since (24a) is of the first 313 order), (25b) and (25d) express, respectively, a homogeneous 314 Robin condition and a Dirichlet condition on J, while (25c) is 315 a Dirichlet condition on p. Finally, the initial conditions read 316 f(R,0) = R, J(R,0) = 1, and p(R,0) = 0, for all  $R \in [0, R_{ext}]$ . 317

#### 4.4 **Discretisation and Results** 318

Let [0,T] be the interval of time over which the system is 319 observed, and let  $0 = t_0 < t_1 < \ldots < t_N = T$  be a partition of 320 [0,T], where  $\tau_n = t_n - t_{n-1}$  is the amplitude of the subinter-321 val  $[t_{n-1}, t_n] \subset [0, T]$ , for n = 1, ..., N, and N is the total num-322 ber of such sub-intervals. Similarly,  $[0, R_{ext}]$  is partitioned as 323  $0 = R_0 < R_1 < \ldots < R_M = R_{\text{ext}}$ , with  $\Delta_m = R_m - R_{m-1}$  being 324 the amplitude of  $[R_{m-1}, R_m] \subset [0, R_{ext}]$ , for  $m = 1, \dots, M$ . Given 325 a generic function q of the radial coordinate and time, the no-326 tation  $q_{m,n} = q(R_m, t_n)$  indicates that q is evaluated at the point 327  $(R_m, t_n)$  of the space-time grid constructed above. 328

Due to the high non-linearity of the system, especially in  $\mathcal{D}$ 329 and A, an explicit Euler method in time is chosen for (24b). To 330

avoid the occurrence of numerical instabilities, the amplitudes  $\Delta_m$  and  $\tau_n$ , which measure, respectively, the increments in space and time, are required to satisfy the constraint  $\Delta_m^2/2\tau_n \leq \mathcal{D}_{ref}$ , for all m = 1, ..., M, and for all n = 1, ..., N, where  $\mathcal{D}_{ref}$  is a constant, referential value of the diffusion coefficient.

The discretised form of (24a)–(24c) is given by

$$f_{m,n} - f_{m-1,n} = \Delta_m \frac{R_m J_{m,n}}{f_{m,n} \lambda_Z(t_n)},$$
(26a)

$$J_{m,n} - J_{m,n-1} = \tau_n \left[ \frac{Q_{m,n-1} - Q_{m-1,n-1}}{\Delta_m} + \frac{Q_{m,n-1}}{R_m} \right], \quad (26b)$$

$$\frac{p_{m,n} - p_{m-1,n}}{\Delta_m} = \frac{(\lambda_{Rm,n})^2}{J_{m,n}k_{m,n}} Q_{m,n},$$
(26c)

with m = 1, ..., M and n = 1, ..., N. For all p = 0, ..., M - 1, and for all q = 0, ..., N,  $Q_{pq}$  is defined as

$$Q_{p,q} = \mathcal{D}_{p,q} \frac{J_{p+1,q} - J_{p,q}}{\Delta_p} - \mathcal{A}_{p,q} J_{p,q}.$$
 (27)

The equations have been implemented independently both in Fortran and in Matlab<sup>©</sup>.

For the simulation of the homogeneous unconfined compression, the parameters specifying the Holmes-Mow permeability defined in (10) are given by  $k_0 = 2.519 \cdot 10^{-3} \text{ mm}^2 \text{MPa}^{-1} \text{s}^{-1}$ , M = 4.638,  $\gamma = 0.0848$ , and  $\phi_{sR} = 0.2$ , while the constants that characterise the Holmes-Mow hyperelastic potential (8) are taken as  $\alpha_0 = 0.11$  MPa,  $\alpha_1 = 0.26$ ,  $\alpha_2 = 0.25$ , and  $\beta = 0.76$ . With the exception of  $\alpha_1$  and  $\alpha_2$ , whose values were assumed, all these data were taken from the experiments on bovine cartilage reported in [5]. The specimen is a cylinder of height H = 2 mmand radius  $R_{\text{ext}} = 3$  mm. Finally, the parameters defining the imposed axial stretch are the target axial displacement  $u_T = 0.4$  mm (corresponding to a final 20% nominal strain) and the time constant  $t_u = 10$  s.

For the whole duration of the simulated experiment, and for the considered set of parameters, only relatively small variations of the volume ratio J (less than the 10%) are observed. Moreover, through most of the (normalised) radius, J remains practically uniform and equal to the initial (undeformed) value of 1, while, close to the lateral boundary, the fluid exudation causes a loss of fluid volume, which is reflected in a decrease in J (Fig. 1). Also the radial component  $P_c^{rR}$  of the constitutive part of the first Piola-Kirchhoff stress tensor (normalised to the material parameter  $\alpha_0$ ) remains virtually uniform through most of the range of the (normalised) radial coordinate, and then decreases to zero to satisfy the boundary condition of zero traction. As time goes on, the stress relaxes because of the exudation of the fluid (Fig. 2).

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**FIGURE 1**. Volume ratio J vs the normalised radial coordinate  $R/R_{\rm ext}$ . The curves are plotted for values of the normalised time  $t/t_u = 0, 1, \dots, 10.$ 



**FIGURE 2**. Radial component  $P_{c}^{rR}$  of the constitutive part of the first Piola-Kirchhoff stress tensor of the solid phase, normalised to the material parameter  $\alpha_0$ , vs the normalised radial coordinate  $R/R_{ext}$ . The curves are plotted for values of the normalised time  $t/t_u = 0, 1, ..., 10$ .

#### 5 Axisymmetric Confined Compression 368

This section focuses on another experimental test that is 369 largely used for characterising the hydraulic and mechanical be-370 haviour of articular cartilage: the axisymmetric confined com-371

pression test. A specimen of tissue is inserted into a cylindrical, 372 impermeable, rigid chamber and compressed by a porous, rigid 373 piston, so that the fluid can escape through it when the speci-374 men is compressed. In the following, confined compression is 375 simulated in displacement control. 376

#### 5.1 Specific Form of the Deformation 377

The solid phase of the tested biphasic medium is isotropic 378 and "transversely homogeneous", i.e., its material properties are 379 allowed to vary only along the axial direction. Furthermore, due 380 to the impermeability of the lower plate and the lateral wall of the chamber, the only non-vanishing component of the fluid velocity 382 is along the symmetry axis of the specimen. In the usual mate-383 rial and spatial cylindrical coordinates  $\{R, \Theta, Z\}$  and  $\{r, \theta, z\}$ , the 384 deformation is given by 385

$$r = \chi^{r}(R, \Theta, Z, t) = R, \qquad (28a)$$

$$\vartheta = \chi^{\vartheta}(R, \Theta, Z, t) = \Theta, \qquad (28b)$$

$$z = \chi^{z}(R, \Theta, Z, t) \equiv \mathfrak{g}(Z, t), \qquad (28c)$$

where  $\chi^z$  has been redefined as a function g of the axial coordi-386 nate Z and time alone. The matrix representing F (from which 387 that of  $C^{-1}$  can be obtained) is 388

$$[F_A^a](Z,t) = \operatorname{diag}[1,1,\lambda_Z(Z,t)], \qquad (29)$$

since the radial and axial stretches are  $\lambda_R = \lambda_{\Theta} = 1$ , while the 389 axial stretch  $\lambda_Z$  satisfies  $J = \lambda_Z = \mathfrak{g}'$ , at all points and all times, 390 with the prime denoting partial differentiation with respect to the 391 axial coordinate Z. 392

#### 5.2 Stress and Balance Equations 393

The form of the deformation specified in (28a)-(28c) im-394 plies that also the matrix representing  $P_c$  is diagonal, and its 395 components can be written as 396

$$P_{\rm c}^{rR}(Z,t) = \tilde{P}_{\rm c}^{rR}(J(Z,t),Z), \qquad (30a)$$

$$P_{\rm c}^{\vartheta\Theta}(Z,t) = \tilde{P}_{\rm c}^{\vartheta\Theta}(J(Z,t),Z), \qquad (30b)$$

$$P_{\rm c}^{zZ}(Z,t) = \tilde{P}_{\rm c}^{zZ}(J(Z,t),Z), \qquad (30c)$$

where the explicit dependence of the constitutive laws on Z has been indicated. Moreover, since all derivatives in directions other than the axial one vanish identically, and since  $P_c^{rR}$  and  $P_c^{\vartheta\Theta}$  are equal to each other, the model equations (12a) and (12b) simplify

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$$\dot{J} = \frac{\partial}{\partial Z} \left[ \frac{k}{J} \frac{\partial P_{\rm c}^{zZ}}{\partial Z} \right], \qquad (31a)$$

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$$\frac{\partial p}{\partial Z} = \frac{\partial P_{\rm c}^{zZ}}{\partial Z}, \qquad (31b)$$

where the permeability k is now assumed to depend explicitly 402 on the axial coordinate. In this formulation, the unknowns are 403 the axial deformation  $\mathfrak{g}$  and the pressure p. If the experiment is 404 performed in displacement control, the following set of boundary 405 and initial conditions must be respected by the unknowns. 406

At the lower boundary (rigid, at rest, and impermeable), 407

$$\mathfrak{g}(0,t) = 0, \qquad (32a)$$

$$\left(\frac{\partial p}{\partial Z}(0,t) = 0 \Rightarrow\right) \frac{\partial P_{\rm c}^{zZ}}{\partial Z}(0,t) = 0.$$
(32b)

At the upper boundary (rigid, moving downward, permeable), 408

$$\mathfrak{g}(H,t) = H - u_T \left[1 - \exp(-t/t_u)\right], \qquad (33a)$$

$$p(H,t) = 0.$$
 (33b) (33b)

Furthermore, the initial conditions are given by g(Z,0) = Z and 409 p(Z,0) = 0, for all  $Z \in [0,H]$ . 410

#### 5.3 "Diffusive Equation" 411

As done in Section 4.3 for the case of the unconfined com-412 pression test, (31a) can be reformulated in the form of a pseudo-413 diffusion-advection equation for the volume ratio J. Indeed, the 414 constitutive definition of  $P_{c}^{zZ}$  leads to the relation 415

$$\frac{\partial P_{\rm c}^{zZ}}{\partial Z} = \frac{\partial \tilde{P}_{\rm c}^{zZ}}{\partial J} \frac{\partial J}{\partial Z} + \frac{\partial \tilde{P}_{\rm c}^{zZ}}{\partial Z} \bigg|_{\rm exp},\tag{34}$$

where the second term on the right-hand-side of (34) denotes 416 the explicit derivative of  $\tilde{P}_{c}^{zZ}$  with respect to the axial coordinate. 417

Thus, by introducing the notation 418

$$\mathcal{D} := \frac{k}{J} \frac{\partial \tilde{P}_{c}^{zZ}}{\partial J}, \qquad (35a)$$

$$-\mathcal{A}J := \frac{k}{J} \frac{\partial \tilde{P}_{c}^{zZ}}{\partial Z} \bigg|_{\exp}, \qquad (35b)$$

and substituting the resulting expressions into (31a), one obtains 419

$$\dot{J} = \frac{\partial}{\partial Z} \left[ \mathcal{D} \frac{\partial J}{\partial Z} - \mathcal{A} J \right].$$
(36)

As for (23),  $\mathcal{D}$  and  $\mathcal{A}$  play the role of the diffusion coefficient and advection velocity, respectively. In this case too, the condition (32b), although written only for  $P_c^{zZ}$ , stems from a condition imposed on the overall axial stress  $P^{zZ} = -p + P_c^{zZ}$ .

Consistently with (35a) and (35b),  $\mathcal{D}$  and  $\mathcal{A}$  can be expressed constitutively as functions of J and Z. As for the unconfined compression test, the approach based on (36) lowers by one the order of the spatial derivatives of g featuring in (31a), but treats J as a free unknown of the model. Therefore, the final form of the model equations reads

$$\frac{\partial \mathfrak{g}}{\partial Z} = J, \qquad (37a)$$

$$\frac{\partial p}{\partial Z} = \frac{\partial \tilde{P}_{c}^{zZ}}{\partial Z},$$
(37b)

$$\dot{J} = \frac{\partial}{\partial Z} \left[ \mathcal{D} \frac{\partial J}{\partial Z} - \mathcal{A}J \right].$$
(37c)

The set (37a)–(37c) comprises three independent scalar equations in the three unknowns g, J and p and is, thus, closed. The boundary conditions must be rephrased compatibly with the new formulation. In the case of a displacement-controlled confined compression test, the boundary conditions become

$$\mathfrak{g}(0,t) = 0, \qquad (38a)$$

$$\left( \mathcal{D} \frac{\partial J}{\partial Z} - \mathcal{A} J \right) \Big|_{Z=0} = 0, \qquad (38b)$$

$$\int_{0}^{H} J(\bar{Z},t) \mathrm{d}\bar{Z} = H - u_{T} [1 - \exp(-t/t_{u})].$$
(38c)

$$p(H,t) = 0, \tag{38d}$$

which have to hold at all times  $t \in [0, T]$ . Equation (38b) is a homogeneous Robin condition on J.

Since (37a) and (37b) are decoupled from (37c), they can be solved *a posteriori*, once J is determined by means of (37c). In particular, it is possible to directly integrate (37a) and (37b), i.e.,

$$\mathfrak{g}(Z,t) = \int_0^Z J(\bar{Z},t) \mathrm{d}\bar{Z}, \qquad (39a)$$

$$p(Z,t) = \tilde{P}_{\rm c}^{zZ}(J(Z,t),Z) - \tilde{P}_{\rm c}^{zZ}(J(H,t),H).$$
(39b)

## 5.4 Discretisation and Results

The numerical solution to (37c) is determined by using central differences for the space derivatives, and an ordinary differential equation (ODE) solver for the time derivatives [12]. The computational domain [0,H] is partitioned as  $0 = Z_1 < \ldots <$  $Z_M = H$ , which determines M - 1 subintervals. In the procedure adopted in this work, all subintervals have the same length

 $\Delta$ . Moreover, for the sake of a lighter notation, the identification 482  $P_c^{zZ} \equiv P$  is made. At the *m*th grid node, with  $m = 2, \dots, (M-1)$ , 448 483 the spatially discretised from of (37c) is given by 449

$$\dot{J}_m = \frac{1}{\Delta^2} \left[ \frac{k_{m+1}}{J_{m+1}} (P_{m+1} - P_m) - \frac{k_m}{J_m} (P_m - P_{m-1}) \right].$$
 (40)

Note that the nodes  $Z_m$ , with m = 2, ..., M - 1, belong to the 450 interior of the computational domain. The values  $J_1$  and  $J_M$ , 451 which correspond to the boundary nodes, must be determined in 452 compliance with the conditions (38b) and (38c). Furthermore, to 453 maintain the second-order-accuracy of the discretisation scheme, 454 a fictitious node  $Z_0 < Z_1$  is introduced, so that the partial deriva-455 tive of J featuring in the Robin condition (38b) can be approx-456 imated by means of the central difference  $(J_2 - J_0)/(2\Delta)$  [12]. 457 The ODEs (40) are then solved in time by using a stable ODE 458 solver, with initial condition  $J_m(0) = 1$ , for all m = 1, ..., M. All 459 numerical simulations have been performed both in Fortran and 460 in Matlab<sup>©</sup>. 461

For the confined compression test,  $P_c^{zZ}$  is given by 462

$$P_{\rm c}^{zZ} = \tilde{P}_{\rm c}^{zZ}(J,Z) = \frac{1}{2}A(Z)\exp\left[(J^2 - 1)\beta\right]\frac{J^2 - 1}{J^{2\beta + 1}},\qquad(41)$$

where  $A = 4 \alpha_0 \beta = 4 \alpha_0 (\alpha_1 + 2\alpha_2)$  [5] is the aggregate elastic 463 modulus (i.e., the stiffness in uni-axial deformation in the linear 464 theory, given by the component L<sup>ZZZZ</sup> of the (material) linear 465 elasticity tensor L), and  $\alpha_0$ ,  $\alpha_1$ ,  $\alpha_2$ ,  $\beta$  are the material constants 466 in the Holmes-Mow hyperelastic potential (8). 467

For this numerical implementation, the undeformed per-468 meability  $k_0$  is obtained by extrapolating the experimental data 469 taken from [10], and the material parameter  $\alpha_0 = A/(4\beta)$  is ob-470 tained from the values of the aggregate modulus A from the ex-471 perimental reported in [11]. Both are expressed by third-order 472 polynomials in the normalised depth Z/H, i.e., 473

$$k_{0}(Z) = \left[-1.4485 \left(\frac{Z}{H}\right)^{3} + 1.4813 \left(\frac{Z}{H}\right)^{2} + 0.0193 \left(\frac{Z}{H}\right) + 0.1371\right] \cdot 10^{-3} \text{mm}^{2} \text{MPa}^{-1} \text{s}^{-1}, \quad (42)$$
  

$$\alpha_{0}(Z) = \left[-1.4953 \left(\frac{Z}{H}\right)^{3} + 3.3255 \left(\frac{Z}{H}\right)^{2} - 2.6711 \left(\frac{Z}{H}\right) + 0.8471\right] \text{MPa}, \quad (43)$$

whereas all other parameters are the same as for the case of un-474 confined compression (M = 4.638,  $\gamma = 0.0848$ , and  $\phi_{sR} = 0.2$ 475 from [5],  $\alpha_1 = 0.26$  and  $\alpha_2 = 0.25$ , whose values are assumed, 476 and  $\beta = \alpha_1 + 2\alpha_2 = 0.76$  from [5]). In this case too, the specimen 477 is a cylinder of initial height H = 2 mm and radius  $R_{\text{ext}} = 3 \text{ mm}$ . 478 The target value and time constant of the imposed axial displace-479 ment are  $u_T = 0.4$  mm (corresponding to a final 20% nominal 480 strain) and  $t_u = 10$  s. 481

Because of the inhomogeneous material properties, the volume ratio J is inhomogeneous through the (normalised) depth of the sample also at stationary state; in particular, the much lower stiffness  $\alpha_0 = A/(4\beta)$  in the superficial zone (close to Z = 1) makes the volumetric compression extreme for the considered overall deformation, with values of  $J \simeq 0.30$  (Fig. 3). Since the pressure p must be zero on the upper boundary, the absolute value of the axial component  $P_{\rm c}^{zZ}$  of the constitutive part of the first Piola-Kirchhoff stress (normalised to the value  $\alpha_0(0)$  that the material parameter  $\alpha_0$  takes at Z = 0 is largest at the upper boundary and equals the absolute value of the total (normalised) stress  $P^{zZ}$ ; at the end of the test, stationary state is practically achieved, as p is zero and consequently  $P_c^{zZ}$  is uniform throughout the tissue depth (Fig. 4).



**FIGURE 3**. Volume ratio J vs the normalised axial coordinate Z/H. The curves are plotted for values of the normalised time  $t/t_{\mu}$  = 0, 1, ..., 10.

#### 6 Discussion

In this work, following the lines of Armstrong et al. [4], who studied unconfined compression of isotropic homogenous cartilage under small deformations, and of Holmes and Mow [5], who studied confined compression of homogenous isotropic cartilage under large deformations, we addressed the unconfined case in the large-deformation setting, and the confined case by removing the hypothesis of homogeneity, thereby allowing some of the material properties to vary along the axis of compression. In both the cases of unconfined and confined compression, we reduced the problem to a diffusion-advection equation in J, which was

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**FIGURE 4.** Axial component  $P_c^{zZ}$  of the constitutive part of the first Piola-Kirchhoff stress tensor of the solid phase, normalised to the value  $\alpha_0(0)$  of the material parameter  $\alpha_0$  at Z = 0, vs the normalised axial coordinate Z/H. The curves are plotted for values of the normalised time  $t/t_u = 0, 1, ..., 10$ .

regarded as the relevant kinematical variable. A result similar to 507 the diffusion-advection equation (36) of the confined case was 508 obtained in [12]. There are, however, two major differences be-509 tween the two approaches. Firstly, the model analysed in [12] 510 was homogeneous and, consequently, could not obtain the ad-511 vection "velocity" A. This "velocity", indeed, arises because of 512 the inhomogeneity of the constitutive law of the axial stress. Sec-513 ondly, in [12], tissue remodelling (an anelastic process) was con-514 sidered, and the hydraulic and mechanical behaviour of the spec-515 imen was studied in the elastic range subsequent remodelling. 516

Although more precise descriptions of articular cartilage 517 have been given [9, 15, 16], where the inhomogeneity and 518 anisotropy of the tissue induced by the presence of the colla-519 gen fibres have been considered, and more general constitutive 520 models can be conceived to include effects such as growth and 521 remodelling (cf., e.g., [17, 18]), the mathematical formulation 522 presented in this work is based on the non-linear biphasic (solid-523 fluid) model. Since we are working with an established theory, 524 and the only "arbitrary" choice is that on the constitutive equa-525 tions, we believe that, by fitting parameters, the vast majority of 526 experimental confined or unconfined tests could validate our nu-527 merical simulations. However, it is clear that homogeneous and 528 isotropic cartilage does not exist and therefore the unconfined 529 case would certainly be a rather artificial fitting of material pa-530 rameters. Moreover, to the best of our knowledge, there is no 531 confined compression test in which both the elastic properties 532

and the permeability have been evaluated. Indeed, the perme-533 ability measurements performed by Maroudas and Bullough [10] 534 do not involve any compression test and, conversely, the com-535 pression tests performed by Schinagl et al. [11] do not involve 536 any permeability measurement. Specifically, the inhomogeneous 537 permeability measurements performed by Maroudas and Bul-538 lough [10] are the only ones we are aware of. Therefore, we 539 cannot infer that our results can have direct experimental valida-540 tion. As far as a comparison with other computational models is 541 concerned, the theoretical derivation of our model has been ob-542 tained by simplifying the theory of biphasic mixtures comprising 543 544 an inviscid fluid and a hyperelastic solid material under the assumption that both phases are incompressible. In this respect, 545 our theoretical results are expected to be consistent with those 546 obtained by the inhomogeneous and anisotropic theory, if the ap-547 propriate model reductions are made. 548

One of the limitations of the method presented here is the isotropy of the material properties. Indeed, mostly due the presence of the collagen fibres, articular cartilage exhibits anisotropic behaviour in both its elastic properties (see, e.g., [9, 16, 19]) and permeability (see, e.g., [8,9,15,16,20]). However, the anisotropy of the tissue was not taken into account here, since the purpose of this work is to show how much information about the mechanical and hydraulic behaviour of the tissue can be extracted also from much simpler models, which do not require elaborated numerical procedures such as the Finite Element Method. Note that neither the fluid inside the cells, nor the intrafibrillar fluid [21,22] are explicitly accounted for in the presented model. Considering these fluids, along with the ions dissolved in them, and their interaction with all the other constituents of the tissue would call for a full electro-chemo-mechanical approach, whose solution would require the employment of sophisticated numerical procedures, especially when large deformations occur. Such a detailed level of modelling is out of the scopes of this paper. The proposed approach is valid also in more complex cases, as long as the further complication is in the non-linearity of the constitutive laws, but ceases to be applicable when the added complication breaks one or more symmetries of the problem. In this case, Finite Element methods often become indispensable.

In our opinion, the advantage of using Finite Differences against Finite Element methods for the problems at hand lies in the fact that the problem reduction shown in the manuscript makes it sufficient to employ one-dimensional grids for solving the model equations in a sufficiently stable, efficient and accurate way, while keeping the computational costs at an acceptable level. This is due to the fact that each of the considered problems is reduced to a set of partial differential equations in which the space dependence appears solely in the partial derivatives with respect to the radial coordinate (in the unconfined compression) or to the axial coordinate (in the confined compression).

The importance of this work is, in fact, in the possibility of testing a given material behaviour (or, more precisely, the

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isotropic version of a material behaviour) in a non-trivial, biphasic, large deformation setting. This means that the result of the
Finite Element implementation of a user-defined material can be
tested against the proposed method, which gives full control on
all physical quantities, since it is based directly on the governing
differential equations.

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