

Mechanics of Cutaneous Wound Rupture

Digendranath Swain^a, Anurag Gupta^{a,*}

^a*Department of Mechanical Engineering,
Indian Institute of Technology,
Kanpur, India, 208016*

Abstract

A cutaneous wound may rupture during healing as a result of stretching in the skin and incompatibility at the wound-skin interface, among other factors. By treating both wound and skin as hyperelastic membranes, and using a biomechanical framework of interfacial growth, we study rupturing as a problem of cavitation in nonlinear elastic materials. We obtain analytical solutions for deformation and residual stress field in the skin-wound configuration while emphasizing the coupling between wound rupture and wrinkling in the skin. The solutions are analyzed in detail for variations in stretching environment, healing condition, and membrane stiffness.

Keywords: Cutaneous wound healing, Wound rupture, Residual stress, Interfacial growth, Wrinkling, Cavitation

1. Introduction

The physiological, cytological, and dehiscence characteristics of cutaneous wound healing are all well researched in the biology literature (Singer and Clark, 1999; Broughton and Rohrich, 2005; Gurtner et al., 2008; Grinnell, 1994; Hahler, 2006; Harhap, 1993). On the other hand, mathematical modelling of wound healing has been conventionally restricted to mostly the biochemical aspects of the problem invoking reaction-diffusion equations for various cellular processes and growth factors (Murray, 2003). Subsequently, however, the importance of mechanical forces and elasticity in restoring the integrity of damaged skin tissues was established (Murray, 2003; Agha et al., 2011; Gurtner et al., 2008; Evans et al., 2013) leading to several proposals of mechanistic models of wound healing (Hall, 2008; Murphy et al., 2011; Tranquillo and Murray, 1992). It has been only recently that cutaneous wound healing is being explored as a problem of biomechanical growth with both skin and the wound modelled as nonlinear elastic materials (Swain and Gupta, 2015; Wu and Amar, 2015; Bowden et al., 2016). The nonlinearity in the elastic response leads to mechanical instabilities in the form of irregular wound geometries (Wu and Amar, 2015), wrinkling in the skin surrounding the wound (Swain and Gupta, 2015) (see also Cerda (2005); Flynn and McCormack (2008); Li and Wang (2011)), and, as shown in the present paper, cavitation in the wound. An understanding of the biomechanics behind such instabilities can provide valuable insights into scar formation and wound management. To the best

*Corresponding author
Email address: ag@iitk.ac.in (Anurag Gupta)

18 of our knowledge, wound cavitation has not been incorporated in any of the previous mathematical
19 models of cutaneous wound closure.

20 The objective of our work is to study the mechanics of wound rupturing within the context of
21 interfacial biological growth and cavitation in hyperelastic membranes. In particular, we are interested
22 to explore the coupling between wound rupture, wrinkling in the unwounded skin, and the residual
23 stress distribution in the skin-wound configuration. Studies in mammalian skin wounds show that
24 the rupture strength of the wounded tissue is less than 10% of the unwounded skin within a week of
25 wounding (Gál et al., 2006; Ramsastry, 2005). This leaves an open skin wound susceptible to rupture
26 under sudden local stretching (Broughton and Rohrich, 2005; Gál et al., 2006). Most importantly, the
27 rupture impairs the healing process and increases the trauma faced by the patient in addition to other
28 health complications (Harhap, 1993). Figure 1(a) shows a ruptured wound surrounded by wrinkled
29 unwounded skin. In our idealized mechanistic model we assume the wound geometry to be circular
30 and consider initiation of rupture to be synonymous with void formation (cavitation) at the center of
31 the wound, see Fig. 1(b). The void appears in the wound at some critical stretching of the skin; it is
32 assumed to be unrelated to other forms of skin cracking such as due to dry weather and old age.

33 In a recent paper, we investigated the emergence of wrinkles and residual stress during wound
34 healing using an interfacial growth model and hyperelastic Varga energies for both wound and skin
35 (Swain and Gupta, 2015). The present work advances on to include the possibility of cavitation, which
36 is tantamount to rupturing, in the wound. In order to do so, we propose a novel two-dimensional
37 (2D) hyperelastic constitutive model for the wound based on a recently developed three-dimensional
38 (3D) hyperelastic strain energy (Xin-Chung and Chang-Jun, 2001). The Varga strain energy density,
39 used previously for the wound, prohibits cavitation and hence cannot be taken suitable for predicting
40 rupture. In fact, cavitation in 2D elastic membranes is restricted by special constitutive requirements
41 (Steigmann, 1992; McMahon et al., 2010; Haughton, 1986). Our model for cavitation in membranes,
42 as a problem of existence and uniqueness of stable bifurcated solutions, follows earlier work in 3D
43 elastic solids (Ball, 1982; Horgan and Polignone, 1995) and 2D hyperelastic membranes (Steigmann,
44 1992; Haughton, 2001, 1990; Haughton and McKay, 1995). The proposed framework can be used to
45 understand the quality of scar formation, post healing, as a consequence of mechanical instabilities
46 emerging from the nonlinear elastic nature of wound and skin. In doing so, it can form a basis for
47 experimentally investigating the precise constitutive nature of the wound, hitherto unestablished in the
48 literature. Our analytical solutions can also provide benchmark results for more sophisticated numerical
49 simulations of elastic instabilities in thin films (Taylor et al., 2014, 2015; Lejeune et al., 2016b,a).

50 In Section 2 we formulate the kinematical structure and the governing equations for the problem
51 at hand. Additionally, we introduce a new 2D hyperelastic strain energy density for the wound and
52 discuss its properties and physical relevance. The boundary value problems for the unwounded skin
53 and the wound are solved in Section 3 to obtain analytical solutions for deformations during wound
54 closure and residual stress distributions. We also discuss criteria for initiation of mechanical instabilities

55 in the form of wrinkling and cavitation. The obtained solutions are discussed in detail in Section 4,
 56 with an emphasis on understanding the effect of wrinkling and stretching of the unwounded skin on
 57 wound rupture and residual stress generation. We briefly discuss the possibility of cavitation at the
 58 intersecting boundary of wound and skin, before concluding our study in Section 5.

59 2. Problem formulation and constitutive assumptions

60 The purpose of this section is to develop a framework which can be used to pursue an analytical study
 61 of biomechanics of rupturing in a wound surrounded by wrinkled skin. We will formulate boundary
 62 value problems, to be solved in the next section, which yield residual stress distribution in wound-skin
 63 configuration and help us analyze the appearance as well as the effects of wrinkling and cavitation.
 64 Towards this end, we consider an instantaneous wound-skin configuration shown as \mathcal{B}_t in Fig. 2, where
 65 a denotes the wound radius and b the outer radius of the skin. We model both wound and skin as 2D
 66 membranes, neglecting their thickness altogether. The deformation is assumed to be axisymmetric so
 67 that the wound-skin arrangement remains circular as wound diameter decreases during healing.

68 2.1. Kinematics and governing equations

69 We provide a quick review of kinematical relations and balance laws required for our further analysis;
 70 details can be seen in our recent work (Swain and Gupta, 2015). The deformations are measured with
 71 respect to a fixed reference configuration \mathcal{B}_0 , see Fig. 2, where the wound and the skin regions are
 72 of radius A and B , respectively, such that $a < A$ to ensure healing. The nature of residual stresses
 73 developed during healing is determined by unloading the wound-skin configuration in \mathcal{B}_t to a zero
 74 stress state. This is achieved, in the present model, by cutting and separating the configuration \mathcal{B}_t
 75 along the wound edge to obtain an intermediate relaxed configuration denoted as \mathcal{B}_i , see Fig. 2. More
 76 quantitative details of this operation are provided later in the section. The assumed axisymmetry of
 77 interfacial growth is manifested in the axisymmetric gap between wound edge and the internal edge of
 78 the skin in \mathcal{B}_i .

79 The position vector in the reference configuration, $\mathbf{X} = R\mathbf{e}_r(\Theta) \in \mathcal{B}_0$ (with polar coordinates
 80 $0 \leq R \leq B$, and $0 \leq \Theta \leq 2\pi$), is related by a continuous bijective map to the position vector in the
 81 current configuration, $\mathbf{x} = r\mathbf{e}_r(\theta) \in \mathcal{B}_t$ (with $0 \leq r \leq b$, and $0 \leq \theta \leq 2\pi$), such that $r = r(R)$ and
 82 $\theta = \Theta$. Here, \mathbf{e}_r and \mathbf{e}_θ are unit basis vectors along radial and circumferential directions of the polar
 83 coordinate system. The corresponding deformation gradient is given by

$$84 \quad \mathbf{F} = r'(R)\mathbf{e}_r \otimes \mathbf{e}_r + \frac{r}{R}\mathbf{e}_\theta \otimes \mathbf{e}_\theta, \quad (1)$$

85 where the superscript prime is used to denote the derivative with respect to R and \otimes stands for
 86 the tensor dyadic product. The nature of healing kinematics is fixed by hypothesising a piecewise
 87 continuous linear map between $\mathbf{X} \in \mathcal{B}_0$ and $\mathbb{X} \in \mathcal{B}_i$ such that $\mathbb{X} = \mathbb{R}(R)\mathbf{e}_r(\Theta)$, where \mathbb{R} is given by
 88 $k_w R$ if $0 \leq R \leq A$ and $k_s R$ if $A \leq R \leq B$. The parameters k_s and k_w are morphoelastic constants

89 for growth of the skin and annihilation of wound, respectively; they are related to mass addition at
 90 the wound-skin interface necessary for wound healing process (Swain and Gupta, 2015). The resulting
 91 growth distortion tensor is clearly isotropic, see Fig. 2, with a form

$$92 \quad \mathbf{F}_g = k_i(\mathbf{e}_r \otimes \mathbf{e}_r + \mathbf{e}_\theta \otimes \mathbf{e}_\theta), \quad (2)$$

93 where k_i should be replaced by k_w and k_s for $0 \leq R \leq A$ and $A \leq R \leq B$, respectively. The elastic
 94 distortion tensor \mathbf{F}_e , satisfying the multiplicative decomposition $\mathbf{F} = \mathbf{F}_e \mathbf{F}_g$ (Rodriguez et al., 1994),
 95 can then be obtained as

$$96 \quad \mathbf{F}_e = \frac{r'(R)}{k_i} \mathbf{e}_r \otimes \mathbf{e}_r + \frac{r}{k_i R} \mathbf{e}_\theta \otimes \mathbf{e}_\theta. \quad (3)$$

97 It is imminent from the above relation that elastic distortion \mathbf{F}_e is incompatible at the wound edge unless
 98 $k_w = k_s$ (Swain and Gupta, 2015). It is this incompatibility that is manifested in the axisymmetric
 99 annular gap between wound and skin regions in \mathcal{B}_i . The incompatibility therefore represents the
 100 differential growth in the two domains. We require $k_s > k_w$ to ensure that the two domains do
 101 not penetrate into each other. Most importantly, the incompatibility is a source for residual stress
 102 distribution in the skin and also has a bearing on the wrinkle formation in the unwounded skin adjacent
 103 to the wound edge (Swain and Gupta, 2015).

104 The stress field in the wound-skin configuration satisfies equation of linear momentum balance
 105 which, for quasi-static deformations, zero body force, and axisymmetry of the problem, yields only one
 106 non-trivial equilibrium condition for stress (Haughton, 2001)

$$107 \quad \frac{dT_{rr}}{dr} + \frac{(T_{rr} - T_{\theta\theta})}{r} = 0 \text{ for } 0 \leq R < A, \quad A < R \leq B \quad (4)$$

108 such that T_{rr} is continuous at $R = A$, where $T_{rr}(R)$, etc. are components of the Cauchy stress with
 109 respect to the polar coordinate system.

110 2.2. Constitutive response of skin

111 The unwounded skin is assumed to behave like a Varga hyperelastic membrane exhibiting non-linear
 112 stress-strain response (Flynn et al., 2011). The anisotropic nature of skin is ignored for analytical
 113 simplicity whereas its viscoelasticity is neglected recognising the slow time scales involved in wound
 114 healing (Hall, 2008). It is justified to model skin as a membrane since the dermis is much softer than the
 115 epidermis allowing for skin to easily slide over the substrate (Cerde, 2005). The strain energy density
 116 (SED) of skin, as considered in the present work, is given by

$$117 \quad W_s(\lambda_1, \lambda_2) = 2\mu_s (\lambda_1 + \lambda_2 + (\lambda_1 \lambda_2)^{-1} - 3), \quad (5)$$

118 where $\mu_s > 0$ can be identified as the shear modulus of the membrane (with dimensions of force/unit length);
 119 λ_1 and λ_2 are principle stretches associated with elastic distortion. The SED in (5), for a Varga hy-
 120 perelastic membrane, supports wrinkling but prohibits cavitation (Steigmann, 1992; Haughton, 2001,
 121 1990; Haughton and McKay, 1995). It is therefore appropriate for the skin but not for the wound.

122 *2.3. Constitutive response of wound*

123 The wound is considered as a solid domain due to its granulating surface acting as a single contractile
 124 body (Broughton and Rohrich, 2005). We ignore the viscoelastic nature of the wound while modelling
 125 it as a hyperelastic membrane whose stress-strain response is less stiffer than skin due to its inferior
 126 properties. The SED function for wound is proposed as

$$127 \quad W_w(\lambda_1, \lambda_2) = C_{01}(\lambda_1 + \lambda_2 - 2) + C_{02}(\lambda_1^{-1} + \lambda_2^{-1} - 2) + C_{03}(\lambda_1\lambda_2 - 1), \quad (6)$$

128 where C_{01} , C_{02} , and C_{03} are material constants whose physical nature will be discussed below. The 2D
 129 SED in (6) is inspired from a SED used to model cavitation in 3D compressible materials (Xin-Chung
 130 and Chang-Jun, 2001). The proposed energy density allows for cavitation in membranes, as shown in
 131 the following section. In rest of this section we analyze it further ensuring that it indeed represents a
 132 physically meaningful energy for hyperelastic membranes.

133 The non-trivial components of the Cauchy stress tensor for an hyperelastic response are given by
 134 (Steigmann, 1992)

$$135 \quad T_{rr} = \frac{1}{\lambda_2} \frac{\partial W_w}{\partial \lambda_1} \quad \text{and} \quad T_{\theta\theta} = \frac{1}{\lambda_1} \frac{\partial W_w}{\partial \lambda_2}. \quad (7)$$

136 Similar expressions can be written for stresses in the skin region. The SED W_w and the derived
 137 components of Cauchy stress must vanish in the stress free configuration, i.e. when $\lambda_1 = \lambda_2 = 1$. The
 138 former of this requirement can be checked by direct substitution in (6). Regarding the latter, we first
 139 obtain the stress components using (7) as

$$140 \quad T_{rr} = \frac{1}{\lambda_2} \left(C_{01} - \frac{C_{02}}{\lambda_1^2} + C_{03}\lambda_2 \right) \quad \text{and} \quad T_{\theta\theta} = \frac{1}{\lambda_1} \left(C_{01} - \frac{C_{02}}{\lambda_2^2} + C_{03}\lambda_1 \right). \quad (8)$$

141 Clearly, the requirement of stress free configuration is satisfied as long as $C_{01} - C_{02} + C_{03} = 0$. More
 142 insight on the nature of the material parameters is obtained by expanding the energy density W_w as a
 143 Taylor series in terms of the Green's strain tensor \mathbf{E} . The leading order terms can then be compared
 144 to the well known linear elastic constants. After retaining only linear and second order coefficients, the
 145 SED can be rewritten as

$$146 \quad W_w = (C_{03}/2)(\text{tr } \mathbf{E})^2 - (1/2)(C_{03} - 2C_{02}) \text{tr } \mathbf{E}^2 + O(\mathbf{E}^3). \quad (9)$$

147 Comparing this with the strain energy density for a plane stress linear elastic problem furnishes $C_{01} =$
 148 $\mu_w(1 - 2\nu)/(1 - \nu)$, $C_{02} = \mu_w/(1 - \nu)$, and $C_{03} = 2\mu_w\nu/(1 - \nu)$, where μ_w is the shear modulus of the
 149 wound material with dimensions of force/unit length and ν is the Poisson's ratio of the material.

150 The principal stretches λ_1 and λ_2 are necessarily positive to prevent disappearance of material. We
 151 require energy W_w to be non-negative for any positive stretch. This is ensured by taking constants C_{01}
 152 and C_{03} to be strictly positive. These restrictions yield $\mu_w > 0$ and $0 < \nu < 1/2$. The energy
 153 density in (6) then has a unique minima at $\lambda_1 = \lambda_2 = 1$. Moreover, the tension-extension inequalities,
 154 $\partial T_{rr}/\partial \lambda_1 > 0$ and $\partial T_{\theta\theta}/\partial \lambda_2 > 0$, are satisfied as long as $C_{02} > 0$. The energy density also satisfies the
 155 Baker-Ericksen Inequality, $(T_{rr} - T_{\theta\theta})(\lambda_1 - \lambda_2) > 0$, for positive C_{01} . The Baker-Ericksen inequality is

156 a well established constitutive restriction for nonlinear elastic materials with important consequences
 157 for the stability and existence of solutions (Truesdell and Noll, 2004). We also note that $W_w \rightarrow \infty$
 158 whenever principal stretches approach ∞ or $0+$. The stresses remain finite in the former limit, but tend
 159 to be unbounded for the latter. Hence, an infinite amount of stress is required to diminish material to
 160 zero volume.

161 We illustrate the behavior of the SED function graphically in Figs. 3(a,b) as a 2D contour map
 162 and a 3D surface plot for a fixed Poisson's ratio ($\nu = 0.3$). The uniaxial and equi-biaxial stress-stretch
 163 responses are shown in Figs. 3(c,d), respectively, for various Poisson's ratios. It can be noted that the
 164 SED has a single minima at $(\lambda_1 = 1, \lambda_2 = 1)$ and the material shows some softening behavior at lower
 165 Poisson's ratios under equi-biaxial stretching.

166 3. Solutions for skin wrinkling and wound cavitation

167 In this section, we construct analytical solutions for deformation and residual stress in skin-wound
 168 configuration. We will first consider the case of unwrinkled skin and then use tension field theory to
 169 derive solutions in the wrinkled region of the skin. Following these we will obtain the solution in the
 170 wound region allowing for the possibility of cavitation at the center of the wound. We will also show
 171 that, beyond a critical bifurcation point, the cavitation solution is always stable and would be preferred
 172 over the homogeneous solution without any cavitation.

173 3.1. Solution for unwrinkled skin

174 The stress fields in the skin can be computed using relations (7), but with SED given by (5), as

$$175 \quad T_{rr}^s = 2\frac{\mu_s}{\lambda_2} \left(1 - \frac{1}{\lambda_1^2 \lambda_2}\right) \quad \text{and} \quad T_{\theta\theta}^s = 2\frac{\mu_s}{\lambda_1} \left(1 - \frac{1}{\lambda_1 \lambda_2^2}\right), \quad (10)$$

176 where the superscript 's' is used to indicate their connection with skin. Substituting these in (4), with
 177 $\lambda_1 = r'/k_s$ and $\lambda_2 = r/(k_s R)$ from (3), we obtain a second order ordinary differential equation (ODE)
 178 $r''rR + (r')^2R - r'r = 0$, which has a straightforward solution $r(R) = \sqrt{C_1 R^2 + C_2}$, with constants C_1
 179 and C_2 to be determined from two boundary conditions. First, we consider $r(B) = b$ to be known, which
 180 is equivalent to prescribing the circumferential stretch at the outer boundary. Second, we also assume
 181 $r(A) = a = \zeta A$ to be given from the healing conditions of the wound, where ζ is the healing constant
 182 (Swain and Gupta, 2015). For a healing wound $\zeta < 1$ and for an atrophic wound $\zeta > 1$. The unknown
 183 constants can then be calculated as $C_1 = (b^2 - a^2)/(B^2 - A^2)$ and $C_2 = (a^2 B^2 - b^2 A^2)/(B^2 - A^2)$.
 184 They can be rewritten in terms of the circumferential stretch at the outer boundary, denoted by λ_{2B} , and
 185 a dimensionless parameter $\alpha = B/A$ as $C_1 = ((\lambda_{2B} k_s \alpha)^2 - \zeta^2)/(\alpha^2 - 1)$ and $C_2 = (\zeta^2 - (\lambda_{2B} k_s)^2) \alpha^2 A^2/(\alpha^2 - 1)$.
 186 The constants are expressed in terms of four physical parameters, ζ , k_s , λ_{2B} , and α , whose prescription
 187 is necessary for the complete solution. The above solution is physically meaningful as long as the asso-
 188 ciated stress fields remain non-negative. Indeed, ideal membranes cannot support compressive stresses
 189 and instead wrinkle to accommodate the compression causing slackness. The solution which allows for
 190 partially wrinkled skin is discussed next.

191 3.2. *Wrinkled skin and tension field theory*

192 We now look for a solution where the skin region, $A \leq R \leq B$, has been partially wrinkled in
 193 a domain, $A \leq R \leq R_c$, adjacent to the wound (see also Swain and Gupta (2015)). The radius R_c
 194 denotes the boundary between wrinkled and unwrinkled skin. The wrinkling will be circumferential in
 195 nature, as shown in Fig. 1(b), if the radial stress remains positive throughout and only the hoop stress,
 196 when calculated in Section 3.1, becomes compressive for $A \leq R \leq R_c$. Our simplistic model cannot
 197 reveal the wavelength of the wrinkling pattern due to vanishing bending rigidity of the membrane. The
 198 solution obtained in the previous section remains valid in the unwrinkled part of the skin, although
 199 with different expressions of C_1 and C_2 . For circumferential wrinkling to appear, the hoop stress $T_{\theta\theta}^s$
 200 must monotonically increase from the inner edge of the skin, where they take a compressive value,
 201 to the outer edge while changing its sign at $R = R_c$. The monotonicity of the hoop stress can be
 202 checked by calculating the gradient, using results in Section 3.1, $(T_{\theta\theta}^s)' = -2\mu_s C_2 k_s / (C_1 R^2 r)$. Note
 203 that, since $\lambda_1 = C_1 R / r k_s$ and $\lambda_1 > 0$, we require $C_1 > 0$. As a result, for circumferential wrinkling
 204 to exist, we should have $C_2 < 0$. Additionally, these constants should be such that T_{rr} remains
 205 positive throughout. The radius R_c can be obtained by solving for R in $T_{\theta\theta} = 0$, which yields a
 206 nonlinear algebraic equation $C_1 \sqrt{C_1 R_c^2 + C_2} = k_s^3 R_c$. Wrinkling can be avoided as long as $R_c \leq A$ or
 207 equivalently when $k_s^3 \geq (b^2 - a^2)\zeta / (B^2 - A^2)$.

208 In order to find the solution in the wrinkled region, $A \leq R \leq R_c$, we use tension field theory as
 209 proposed by Pipkin and Steigmann (Pipkin, 1986; Steigmann, 1990). The essential idea is to regularise
 210 the original energy to obtain a relaxed energy which is compatible with the wrinkled solution. The
 211 ‘natural width’ $n(\lambda_1)$ of the membrane is given by λ_2 which can be solved in terms of λ_1 using $T_{\theta\theta}^s = 0$
 212 to obtain $\lambda_2 = 1/\sqrt{\lambda_1}$ (Pipkin, 1986; Haughton and McKay, 1995). The relaxed SED function, denoted
 213 by W_s^* , is defined as

$$214 \quad W_s^*(\lambda_1) = W_s(\lambda_1, n(\lambda_1)) = 2\mu_s \left(\lambda_1 + 2/\sqrt{\lambda_1} - 3 \right), \quad (11)$$

215 where we have used superscript $*$ to indicate the tension field variables. The governing equation
 216 (4) simplifies to $dT_{rr}^*/dr + T_{rr}^*/r = 0$ yielding $rT_{rr}^* = \text{const.}$, which in conjunction with (10)₁ gives
 217 $R(1 - \lambda_1^{-1.5}) = \text{const.}$ This can be integrated to calculate the deformation in the wrinkled region as
 218 (Haughton and McKay, 1995; Swain and Gupta, 2015)

$$219 \quad r^*(R) = \int \left(\frac{R}{R - \delta_1} \right)^{2/3} k_s dR + \delta_2, \quad (12)$$

220 where δ_1 and δ_2 are constants. The four unknown constants, δ_1, δ_2 in the wrinkled solution and C_1, C_2
 221 in the unwrinkled solution, can be determined from two boundary conditions given by the continuity of
 222 deformation and radial stress at wrinkle boundary $R = R_c$ apart from two other boundary conditions
 223 prescribing displacements at the inner and the outer edge of the skin domain, as in Section 3.1.

224 *3.3. Solution allowing for cavitation in the wound*

225 The governing equation for deformation in the wound region can be obtained by substituting stresses
 226 from (8) into (4), and using $\lambda_1 = r'/k_w$ and $\lambda_2 = r/(k_w R)$, as

$$227 \quad 2r^2 Rr'' - r^2 r' + R^2 (r')^3 = 0. \quad (13)$$

228 To solve the preceding second order ODE, we need two boundary conditions. Towards this end, we
 229 prescribe displacement at the wound edge $r(A) = k_w \lambda A$, where λ is the applied stretch transmitted
 230 through the skin (or equivalently through the wound edge stresses). The second boundary condition
 231 is given by $r(0) = 0$ when no cavity appears at the center of the wound domain, or $r(0^+) = \gamma > 0$
 232 when a cavity (or void) appears at the center. In the latter scenario, the surface of the cavity must
 233 be traction free, i.e. $T_{rr}(0^+) = 0$ (Steigmann, 1992). A solution with homogeneous deformation
 234 of the kind $r = k_w \lambda R$ satisfies (13) and the boundary conditions without cavity. We are however
 235 interested in finding a solution which allows for cavity. To do so, we introduce $\beta(R) = \lambda_1/\lambda_2 = Rr'/r$
 236 and rewrite (13) as $2R\beta' + \beta(\beta - 1)(\beta + 3) = 0$. The first order ODE can be integrated to obtain
 237 $R(\beta) = D\beta^{2/3}(\beta + 3)^{-1/6}/\sqrt{\beta - 1}$ and $r(\beta) = C\sqrt{\beta + 3}/\sqrt{\beta - 1}$, where D and C are constants of
 238 integration, to be determined using $R(\beta_A) = A$ and $r(\beta_A) = k_w \lambda A$, where $\beta_A = \beta(A)$. After solving
 239 for these constants we obtain

$$240 \quad R(\beta) = A \left(\frac{\beta}{\beta_A} \right)^{2/3} \left(\frac{\beta_A + 3}{\beta + 3} \right)^{1/6} \left(\frac{\beta_A - 1}{\beta - 1} \right)^{1/2} \quad \text{and} \quad (14)$$

$$241 \quad r(\beta) = k_w \lambda A \left(\frac{\beta + 3}{\beta_A + 3} \right)^{1/2} \left(\frac{\beta_A - 1}{\beta - 1} \right)^{1/2}. \quad (15)$$

242 The parameter β can be eliminated between these two expression to find the deformation $r(R)$ in the
 243 wound. The constant β_A will be determined below. Due to the positivity of the principal stretches,
 244 $\beta > 0$. Also, for finite $r(R = 0^+)$, $\beta(0^+) = 0$. Hence, for the cavitation solution, $0 < \beta < 1$, since
 245 otherwise $\beta' < 0$ which leads to a contradictory result. This also necessitates $0 < \beta_A < 1$. At the
 246 center of the wound (15) yields

$$247 \quad \gamma = k_w \lambda A \sqrt{3(1 - \beta_A)}/\sqrt{\beta_A + 3}. \quad (16)$$

248 We can study the cavitation phenomenon as a bifurcation problem. Indeed, there is a critical value
 249 of the stretch λ , controlled at the wound edge, at which the cavitation solution exists; the critical value,
 250 denoted as λ_c , will be calculated below. For $\lambda < \lambda_c$, only the homogeneous solution (without cavity)
 251 is possible and there is no solution which allows for a cavity at the center of the wound. On the other
 252 hand, for $\lambda > \lambda_c$ it is possible to obtain another solution which allows for cavitation. We will identify
 253 $\lambda = \lambda_c$ as the critical point for bifurcation. As we shall see later in the section, the cavitation solution
 254 is energetically stable and will therefore be preferred over the homogeneous solution beyond the critical
 255 point. In order to find λ_c , we begin by noting that $\gamma = 0$ at the critical point of bifurcation, which
 256 can be used to calculate the critical value of β_A as $\beta_{Ac} = 1$. The critical stretch λ_c will be obtained
 257 using the stress free boundary condition at $R = 0$ (i.e. on the edge of cavity). First, using (14) and

(15), we write principle stretches as $\lambda_2 = \lambda(\beta_A/\beta)^{2/3}(\beta + 3)^{2/3}(\beta_A + 3)^{-2/3}$ and $\lambda_1 = \beta\lambda_2$, which are then substituted into the expression for radial stress in (8)₁. The traction free boundary condition $T_{rr}(0) = 0$, on using $\beta = 0$ at $R=0$, immediately yields $-C_{02}(3 + \beta_A)^2 + 9\lambda^3\beta_A^2C_{03} = 0$. Recalling that $C_{03} = 2\nu C_{02}$, we can solve this equation to obtain a formula for β_A as $\beta_A = 3/\left(3\sqrt{2\nu\lambda^3} - 1\right)$. Furthermore, using $\beta_{Ac} = 1$ gives the critical value of stretch, $\lambda_c = 2(9\nu)^{-1/3}$. Interestingly, the critical stretch depends only on the Poisson's ratio. The derived relation for β_A can be used in (16) to get an expression for the size of the cavity,

$$\gamma = k_w\lambda A\sqrt{1 - 4/(3\sqrt{2\nu\lambda^3})}. \quad (17)$$

The variation in β_A with respect to applied stretch for various Poisson's ratios is shown in Fig. 4(a). The intersection of $\beta_A = 1$ line with the plotted curves provide the critical stretch for the respective Poisson's ratio. For any stretch applied beyond these points, β_A diminishes non-linearly. In Fig. 4(b) we illustrate void growth with respect to the applied stretch. Clearly, the growth, as well as the critical stretch at which growth initiates, varies with Poisson's ratio of the wound material.

Finally, we verify whether cavitation is energetically stable for edge stretch magnitudes beyond the critical value. The total stored energy of the axisymmetrically deforming circular wound is given by $E = \int_0^A 2\pi W_w R dR$. Its evaluation is greatly simplified by noting an identity, $2RW_w = (R^2(W_w - (\lambda_1 - \lambda_2)\partial W_w/\partial\lambda_1))'$, which can be verified using (3), (4), and (7). As a result, the total energy reduces to $E = \pi A^2(W_w - (\lambda_1 - \lambda_2)\partial W_w/\partial\lambda_1)$, where all the fields are evaluated at $R = A$. Using the energy density W_w as postulated in (6), we first calculate the total energy for the homogeneous solution, i.e. when $\lambda_1 = \lambda_2 = \lambda$, as $E_{hom} = \pi A^2(\lambda - 1)^2(2C_{02}/\lambda + C_{03})$. Analogously, we can obtain the total energy for the non-homogenous cavitation solution, where at the wound edge $\lambda_1 = \beta_A\lambda$ and $\lambda_2 = \lambda$, as

$$E_{nonhom} = \pi A^2 \left((\lambda - 1)^2 C_{03} + C_{02} \left(2(\lambda - 2) + \frac{2\beta_A + \beta_A^2 - 1}{\lambda\beta_A^2} \right) \right). \quad (18)$$

The difference $E_{hom} - E_{nonhom} = \pi A^2 C_{02}(\beta_A - 1)^2/\lambda\beta_A^2$ is always positive since $\lambda > 0$ and $C_{02} > 0$. Therefore the cavitation solution is energetically favourable over the homogeneous solution.

4. Discussion

We are broadly interested in three phenomena during rupture of a cutaneous wound: wrinkle formation in the skin adjacent to the wound, emergence of residual stresses in the skin due to wound healing and wound edge incompatibility, and void formation in the wound as a result of cavitation in the wound membrane. The hyperelastic membrane models for skin and wound, as proposed in the previous section, allow for these occurrences. We will now discuss in detail the physical nature of our model as well as the solutions, and study their dependence on various parameters. To begin with, we plot (in Fig. 5) radial and hoop stresses derived from the Varga model of skin given in (5). We fix $\alpha = B/A = 3$ and obtain stress distributions for varying incompatibility parameter k_s , healing constant ζ , and applied stretch λ_{2B} . The skin domain is stress free when all the parameters are unity. For all

292 other considered values, the circumferential stress takes negative values in a finite region, indicating the
 293 emergence of wrinkling. The stress fields in the wrinkled region need to be modified using tension field
 294 theory as shown in a previous section. Clearly, deviation from unity in the value of k_s , ζ , or λ_{2B} leads
 295 to residual stresses, more so when the deviation is in more than one parameter. Most importantly,
 296 these results show how the stress fields in the skin could potentially change as a result of wound healing
 297 (Swain and Gupta, 2015). This is expected since, while healing is in progress, the cellular processes at
 298 the skin edge exert internal forces which are primarily responsible for such changes. Moreover, when
 299 applied stretch is considered (blue lines), the magnitude of residual stresses is the highest whereas the
 300 wrinkling radius minimum. It should be noted that we have ignored the natural pre-tension of the
 301 skin while plotting these stresses, which should otherwise be superimposed with the obtained residual
 302 stresses to find the total stress distribution. In the following subsections we further elaborate the role
 303 of various parameters on wrinkling, wound edge stresses, rupture of wound, and critical stretch for void
 304 formation.

305 *4.1. Role of applied stretch at the outer edge of the skin*

306 The stretching of skin can occur due to normal motion of the body (of various joints, muscles, and
 307 limb), change in postures, or even due to respiration and neck movements. It could be severe if skin
 308 rubs along with external objects or if the body experiences sudden motion as in sports. The severity
 309 of stretching is captured by the variable λ_{2B} in our model. The effects of stretching on wrinkling,
 310 stress distribution, and cavitation are summarized in Fig. 6 and Table 1(a). The dashed lines in Fig. 6
 311 are the tension field solutions while the solid lines are obtained without incorporating tension field.
 312 Expectedly, wrinkling decreases the magnitude of radial stresses due to the lateral slackening. Our
 313 results also show that the wrinkled region diminishes with increased radial stretching. This is due to
 314 increased tensile stresses in the skin owing to higher boundary stretching. The stresses at the wound
 315 edge are also proportional to the applied stretch. An increase in applied stretch may therefore lead
 316 to sudden void formation and subsequent rupturing in wounds whose shear modulus is less than 0.9
 317 times the modulus of skin. In other words, severe stretching will always lead to wound rupture. Lower
 318 values of stretching however can rupture wounds only up to specific Poisson's ratios, as shown in Table
 319 1(a). The critical stretch behavior in Table 1(a) shows that even small amount of stretching inside the
 320 wound is sufficient for rupture whenever the externally applied stretches are large.

321 *4.2. Role of the location of applied stretch*

322 The applied stretch is provided at the outer edge of the skin whose location is fixed by the parameter
 323 α for a given A . The considered location may vary depending on the physical position of the wound
 324 on the body, for example wounds on knee or elbow joints are subjected to local stretching whereas
 325 wounds on chest and abdominal joints are exposed to only far field stretching. Moreover, sports related
 326 trauma in the wound can occur due to arbitrary contact of skin with external objects thereby inducing
 327 in-plane stretching. The effects of α on wrinkle characteristics, stress distribution, and cavitation are

328 summarized in Fig. 7 and Table 1(b). A localized stretching results in smaller wrinkles due to increased
 329 stresses in the skin and at the wound edge. Moreover, localized stretching can rupture all wounds with
 330 shear moduli 0.7 times less than that of skin. When the location of applied stretch moves away from
 331 the wound the rupturing becomes restricted to specific Poisson's ratios. The critical stretch required
 332 for cavitation increases with an increase in the radius of the outer edge of the skin.

333 4.3. Role of healing constant

334 The parameter ζ represents the nature of wound healing; $\zeta < 1$ implies that the wound is undergoing
 335 healing and $\zeta > 1$ denotes an atrophic condition of the wound which could be due to nutritional
 336 deficiency, hygiene, or infection. Smaller values of ζ (below unity) indicate faster healing, for example
 337 $\zeta = 0.95$ (change in radius is 5%) implies faster contraction than $\zeta = 0.99$ (change in radius is 1%). The
 338 healing of the wound can be hastened with the help of appropriate wound treatment and proper hygiene.
 339 The effects of healing condition on wrinkling in the skin, residual stresses, and rupture behavior of the
 340 wound are reported in Fig. 8 and Table 1(c). It is seen that the wounds which heal faster create larger
 341 wrinkles around the wound. This is due to relatively larger stresses in the skin and at the wound edge.
 342 The observed wrinkling behavior agrees well with the existing literature (Cerdeira, 2005; Geminard et al.,
 343 2004; Swain and Gupta, 2015; Flynn and McCormack, 2008). With faster healing, say $\zeta = 0.95$, a wider
 344 range of wounds with Poisson's ratios upto 0.172 can be ruptured. However, when the healing is slow, or
 345 the wound is in atrophic condition, the cavitation can occur in wounds only for a very restricted range
 346 of material parameters. Clearly, the wounds in atrophy need larger stretching to rupture. Moreover,
 347 the critical stretch required for rupturing increases with dilapidated healing condition of the wound.

348 4.4. Role of incompatibility at the wound edge

349 The incompatibility at the wound edge controls the morphoelastic behavior of cutaneous wound
 350 closure and is directly related to generation of residual stresses in the wound-skin arrangement. In our
 351 model, the incompatibility is controlled by the difference $k_s - k_w$. In the present discussion we report
 352 the effect of k_s variation on various aspect of wound healing. The parameter k_s represents the cellular
 353 action near the wound edge on the skin side leading the skin to grow towards the wound center to
 354 achieve healing (Swain and Gupta, 2015). A large value of k_s represents higher level of cell production
 355 in the skin side of the wound edge. The wrinkling behavior, stresses, and rupture behavior for various
 356 k_s values can be seen in Fig. 9 and Table 1(d). The wrinkling radius increases marginally with an
 357 increase in k_s , leading to higher stresses. See also Fig. 5, where it is clear that k_s influences both
 358 wrinkling behavior and stresses as a result of inhomogeneous expansion. A sufficiently high value of
 359 k_s can rupture all wounds with shear modulus less than 0.7 times the skin modulus due to high values
 360 of wound edge stress. For smaller k_s values wound rupture is possible only with limited constitutive
 361 conditions. The critical stretch required for void formation decreases with k_s .

Parameter	Parameter Values	Important Parameters in Skin and Wound			
		Wrinkle radius R_c/A	Wound edge Stress T_{rr}/μ_s	Poisson's ratio ν	Critical stretch λ_c
(a) Role of applied stretch at the outer edge of skin ($B = 3A$, $\zeta = 0.95$, $k_s = 1.1$, and $\mu_w = 0.7\mu_s$)					
λ_{2B}	1.01	1.41	0.448	0.172	1.729
	1.02	1.24	0.545	0.308	1.423
	1.05	1.02	0.790	$\mu_w/\mu_s \leq 0.9$	
(b) Role of the location of the applied stretch ($\lambda_{2B} = 1.01$, $\zeta = 0.95$, $k_s = 1.1$, and $\mu_w = 0.7\mu_s$)					
$\alpha = B/A$	2	1.10	0.673	$\mu_w/\mu_s \leq 0.7$	
	3	1.41	0.448	0.172	1.729
	4	1.62	0.370	0.101	2.065
(c) Role of the healing constant ($B = 3A$, $\lambda_{2B} = 1.01$, $k_s = 1.1$, and $\mu_w = 0.7\mu_s$)					
ζ	0.95	1.41	0.448	0.172	1.729
	0.99	1.33	0.360	0.094	2.113
	1.01	1.28	0.315	0.065	2.389
(d) Role of the wound edge incompatibility ($B = 3A$, $\lambda_{2B} = 1.01$, $\zeta = 0.9$, and $\mu_w = 0.7\mu_s$)					
k_s	1.05	1.42	0.463	0.189	1.676
	1.10	1.48	0.552	0.321	1.404
	1.15	1.52	0.631	$\mu_w/\mu_s \leq 0.7$	
(e) Role of the elasticity of the wound ($B = 3A$, $\lambda_{2B} = 1.01$, $\zeta = 0.95$, and $k_s = 1.1$)					
μ_w/μ_s	0.65	1.41	0.448	0.212	1.611
	0.70	1.41	0.448	0.172	1.729
	0.75	1.41	0.448	0.142	1.844

Table 1: The effect of different parameters on wrinkling radius R_c/A , wound edge radial stresses T_{rr}/μ_s , critical stretch λ_c , and maximum allowable wound Poisson's ratio ν .

362 4.5. Role of elasticity of the wound

363 The elastic characteristic of the wound can be improved by means of medical treatment. In any
364 case, the ultimate tensile strength of a wound remains much inferior than the skin until full healing
365 is achieved (Ramsastry, 2005; Gál et al., 2006). The effect of wound shear modulus on the rupture
366 characteristics is shown in Table 1(e). It is clear that both wrinkling behaviour and stress distribution

367 remain invariant for all the cases studied. A variation in relative stiffness is therefore seen to effect
 368 only the rupturing of the wound. As shown in the table, an improvement in wound properties can
 369 restrict rupturing for a wide range of Poisson's ratio. Wounds with worse properties will of course
 370 rupture easily for a large class of wounds. The critical stretch required for void formation increases on
 371 improving the wound elasticity. Hence, stiffer wounds may not allow for rapid rupture.

372 4.6. Possibility of cavitation at the wound-skin interface

373 In Section 3.3, and the subsequent discussion, the cavitation in the circular wound membrane has
 374 been assumed to take place at the center. This leads to an axisymmetrical problem with a straightfor-
 375 ward analytical solution. Another possibility is to look for solutions with cavitation at the interface of
 376 wound and skin. This would however result in a non-axisymmetric problem without analytical solutions.
 377 In this section, we nevertheless visit this scenario under some simplifying assumptions while restricting
 378 ourselves to only energy based arguments. For our analysis, we consider a circular disc of radius A , such
 379 that one half of the disc ($0 \leq R \leq A$, $0 \leq \theta < \pi$) is occupied by the wound membrane and the other half
 380 ($0 \leq R \leq A$, $\pi \leq \theta < 2\pi$) by the skin membrane. We look for the possibility of cavitation at the center
 381 of this disc at the interface of wound and skin membranes. The solution to the resulting problem is as-
 382 sumed to remain axisymmetric. The total stored energy of the disc containing the wound-skin interface
 383 can be evaluated using $E = \int_0^A \pi(W_w + W_s)RdR$ which, following the procedure used in Section 3.3, can
 384 be written as $E = (\pi R_o^2/2)((W_w + W_s) - (\lambda_1 - \lambda_2)\partial(W_w + W_s)/\partial\lambda_1)$, where all the fields are evalu-
 385 ated at $R = A$. As before, the homogeneous solution is given by $\lambda_1 = \lambda_2 = \lambda$ and the non-homogeneous
 386 cavitation solution by $\lambda_1 = \beta_A \lambda_2 = \beta_A \lambda$. The difference in stored energies due to homogeneous and non-
 387 homogeneous solution can then be estimated as $\Delta E = (\pi R_o^2/2)(C_{02}\lambda + 2\mu_s)(\beta - 1)^2/(\beta\lambda)^2$. Clearly,
 388 the non-homogeneous solution with the wound-skin interface is stable and has a lower energy than the
 389 homogeneous solution. However, since Varga membrane does not support cavitation, the cavitation
 390 will occur only in the wound side of the disc.

391 5. Concluding remarks

392 We have revisited the problem of cutaneous wound healing by incorporating wound rupture in a
 393 recently developed framework of interfacial biomechanical growth (Swain and Gupta, 2015). We pro-
 394 posed a novel hyperelastic strain energy to model the 2D wound membrane which allows for cavitation,
 395 unlike previously employed Varga membranes. The resulting framework predicted simultaneous oc-
 396 currence of ruptured wound and wrinkled skin in a region adjacent to the wound edge. The relevant
 397 boundary value problems were solved analytically and closed form solutions obtained for deformation
 398 and stresses in skin-wound configuration. Both wrinkling and cavitation emerged as stable solutions to
 399 the bifurcation problems in nonlinear elasticity of 2D membranes. The present work can be advanced in
 400 several directions. Most importantly, experimental investigations on the constitutive nature of wound
 401 membrane can provide data for verification of the proposed hyperelastic model. On the other hand,

402 precise experiments for measuring the residual stress distribution in the skin during wound healing
403 can be used to fit unknown parameters in our model making it useful for practical biomedical appli-
404 cations. The theoretical framework of the model can also be improved, although at the cost of losing
405 analytical solvability, by including membrane curvature, non-circular wounds, finite bending rigidity,
406 and viscoelasticity, among other considerations.

407 **Conflict of interest statement**

408 The authors declared no potential conflicts of interest with respect to the research, authorship,
409 and/or publication of this article.

410 **Funding**

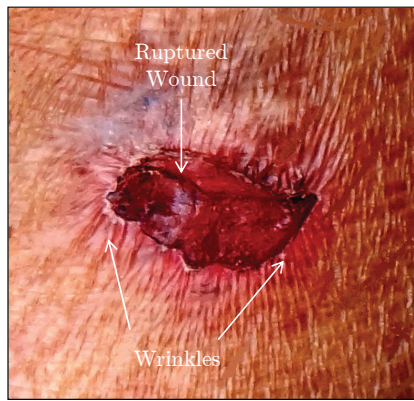
411 This research received no specific grant from any funding agency in the public, commercial, or
412 not-for-profit sectors.

413 **References**

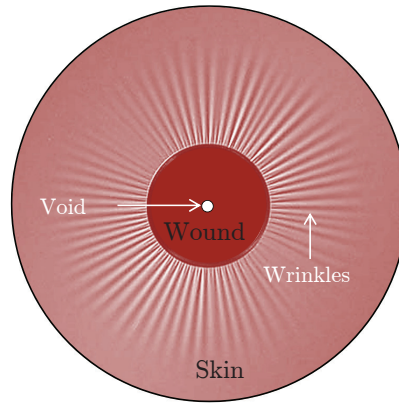
- 414 Agha, R., Ogawa, R., Pietramaggiore, G., Orgill, D.P., 2011. A review of the role of mechanical forces
415 in cutaneous wound healing. *J. Surg. Res.* 171, 700–708.
- 416 Ball, J.M., 1982. Discontinuous equilibrium solutions and cavitation in nonlinear elasticity. *Philos. T.*
417 *Roy. Soc. A* 306, 557–611.
- 418 Bowden, L.G., Byrne, H.M., Maini, P.K., Moulton, D.E., 2016. A morphoelastic model for dermal
419 wound closure. *Biomech. Model Mechanobiol.* 15, 663–681.
- 420 Broughton, G., Rohrich, R.R., 2005. Wounds and scars. *Sel. Readings Plast. Surg.* 10, 1–54.
- 421 Cerda, E., 2005. Mechanics of scars. *J. Biomech.* 38, 1598–1603.
- 422 Evans, N.D., Oreffo, R.O.C., Healy, E., Thurner, P.J., Man, Y.H., 2013. Epithelial mechanobiology,
423 skin wound healing, and stem cell niche. *J. Mech. Behav. Biomed. Mater.* 28, 397–409.
- 424 Flynn, C., McCormack, B.A.O., 2008. A simplified model of scar contraction. *J. Biomech.* 41, 1582–
425 1589.
- 426 Flynn, C., Taberner, A., Nielsen, P., 2011. Modeling the mechanical response of in vivo human skin
427 under rich set of deformations. *Ann. Biomed. Eng.* 39, 1935–1946.
- 428 Gál, P., Toporcer, T., Vidinský, B., Mokřý, M., Novotný, M., Kilík, R., Jr., K.S., Gál, T., Sabo, J.,
429 2006. Early changes in the tensile strength and morphology of primary sutured skin wounds in rats.
430 *Folia Biol. (Praha)* 52, 109–115.
- 431 Geminard, J.C., Bernal, R., Melo, F., 2004. Wrinkle formations in axi-symmetrically stretched mem-
432 branes. *Eur. Phys. J. E Soft Matter* 15, 117–126.

- 433 Grinnell, F., 1994. Fibroblasts, myofibroblasts, and wound contraction. *J. Cell Biol.* 124, 401–404.
- 434 Gurtner, G.C., Werner, S., Barrandon, Y., Longaker, M.T., 2008. Wound repair and regeneration.
435 *Nature* 453, 314–321.
- 436 Hahler, B., 2006. Surgical wound dehiscence. *Medsurg Nurs.* 15, 296–300.
- 437 Hall, C.L., 2008. Modeling of Some Biological Materials using Continuum Mechanics. Ph.D. thesis.
438 Queensland University of Technology, Australia.
- 439 Harhap, M. (Ed.), 1993. Complications of Dermatologic Surgery. Springer-Verlag Berlin.
- 440 Haughton, D.M., 1986. On non-existence of cavitation in incompressible elastic membranes. *Q. J.*
441 *Mech. Appl. Math.* 39, 289–296.
- 442 Haughton, D.M., 1990. Cavitation in compressible elastic membranes. *Int. J. Eng. Sci.* 28, 163–168.
- 443 Haughton, D.M., 2001. Elastic membranes, in: Fu, Y.B., Ogden, R.W. (Eds.), *Non-linear Elasticity:*
444 *Theory and Applications.* Cambridge University Press, pp. 233–267.
- 445 Haughton, D.M., McKay, B.A., 1995. Wrinkling of annular discs subjected to radial displacements.
446 *Int. J. Eng. Sci.* 33, 335–350.
- 447 Horgan, C.O., Polignone, D.A., 1995. Cavitation in nonlinearly elastic solids: A review. *ASME Appl.*
448 *Mech. Rev.* 48, 471–485.
- 449 Lejeune, E., Javili, A., Linder, C., 2016a. An algorithmic approach to multi-layer wrinkling. *Extreme*
450 *Mech. Lett.* 7, 10–17.
- 451 Lejeune, E., Javili, A., Linder, C., 2016b. Understanding geometric instabilities in thin films via a
452 multi-layer model. *Soft Matter* 12, 806–816.
- 453 Li, B., Wang, J.H.C., 2011. Fibroblasts and myofibroblasts in wound healing: Force generation and
454 measurement. *J. Tissue Viability* 20, 108–120.
- 455 McMahon, J., Goriley, A., Tabor, M., 2010. Spontaneous cavitation in growing elastic membranes.
456 *Math. Mech. Solids* 15, 57–77.
- 457 Murphy, K.E., Hall, C.L., McCue, S.W., McElwain, D.L.S., 2011. A two-compartment mechanochemical
458 model of the roles of transforming growth factor β and tissue tension in dermal wound healing. *J.*
459 *Theor. Biol.* 272, 145–159.
- 460 Murray, J.D., 2003. *Mathematical Biology II: Spatial Models and Biomedical Applications.* Springer-
461 Verlag, New York.
- 462 Pipkin, A.C., 1986. The relaxed energy density for isotropic elastic membranes. *IMA J. Appl. Math.*
463 36, 85–99.

- 464 Ramsastry, S.S., 2005. Acute wounds. *Clin. Plast. Surg.* 32, 195–208.
- 465 Rodriguez, E.K., Hoger, A., McCulloch, A.D., 1994. Stress-dependent finite growth in soft elastic
466 tissues. *J. Biomech.* 21, 455–467.
- 467 Singer, A.J., Clark, R.A.F., 1999. Cutaneous wound healing. *New Engl. J. Med.* 341, 738–746.
- 468 Steigmann, D.J., 1990. Tension-field theory. *Proc. Roy. Soc. Lond. A Mat.* 429, 141–173.
- 469 Steigmann, D.J., 1992. Cavitation in elastic membranes – an example. *J. Elasticity* 28, 277–187.
- 470 Swain, D., Gupta, A., 2015. Interfacial biological growth during closure of a cutaneous wound: Stress
471 generation and wrinkle formation. *Soft Matter* 11, 6499–6508.
- 472 Taylor, M., Bertoldi, K., Steigmann, D.J., 2014. Spatial resolution of wrinkle patterns in thin elastic
473 sheets at finite strain. *J. Mech. Phys. Solids* 62, 163–180.
- 474 Taylor, M., Davidovitch, B., Qiu, Z., Bertoldi, K., 2015. A comparative analysis of numerical approaches
475 to the mechanics of elastic sheets. *J. Mech. Phys. Solids* 79, 92–107.
- 476 Tranquillo, R.T., Murray, J.D., 1992. Continuum model of fibroblast-driven wound contraction:
477 inflammation-mediation. *J. Theor. Biol.* 158, 135–172.
- 478 Truesdell, C., Noll, W., 2004. *The Non-Linear Field Theories of Mechanics*, Third Edition. Springer-
479 Verlag Berlin Heidelberg.
- 480 Wu, M., Amar, M.B., 2015. Growth and remodelling for profound circular wounds in skin. *Biomech.*
481 *Model Mechanobiol.* 14, 357–370.
- 482 Xin-Chung, S., Chang-Jun, C., 2001. Exact solution for cavitated bifurcation for compressible hypere-
483 lastic materials. *Int. J. Eng. Sci.* 39, 1101–1117.



(a)



(b)

Figure 1: (a) Photograph of a ruptured wound near the knee joint surrounded by wrinkled skin. (b) Idealized representation of a ruptured wound as considered in the present work with cavitation at the center of the wound while being surrounded by an axisymmetric distribution of wrinkling in the unwounded skin.

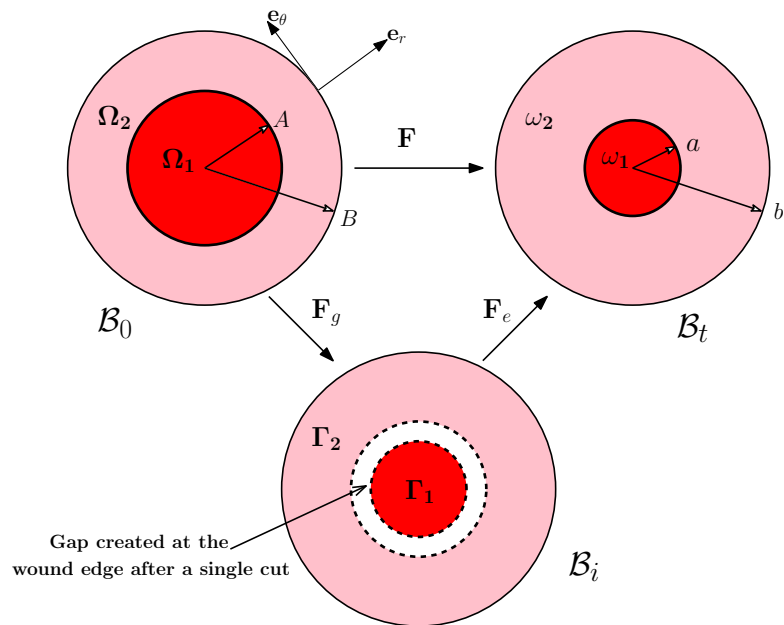
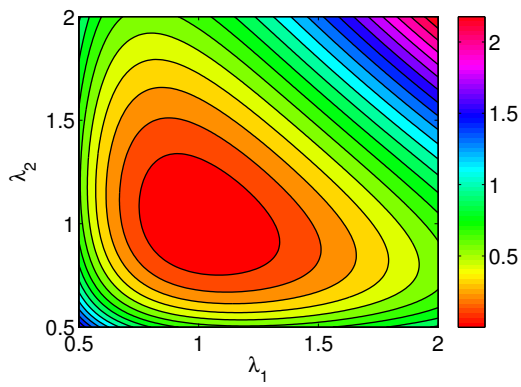
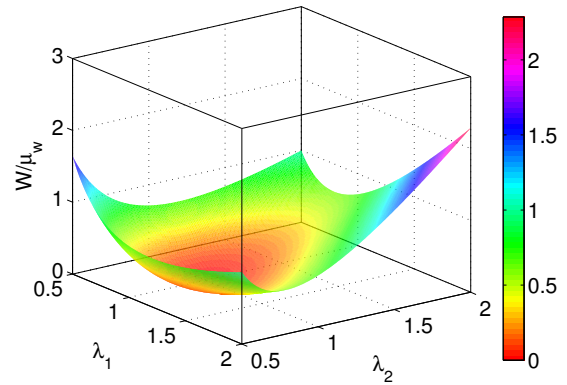


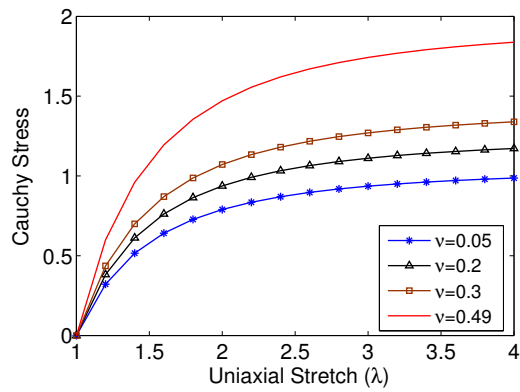
Figure 2: Kinematics of wound healing, where \mathcal{B}_0 is the reference configuration with Ω_1 (wound) and Ω_2 (skin), \mathcal{B}_i is the intermediate stress free configuration with Γ_1 (wound) and Γ_2 (skin), and \mathcal{B}_t is the current configuration with ω_1 (wound) and ω_2 (skin). Figure adapted from (Swain and Gupta, 2015).



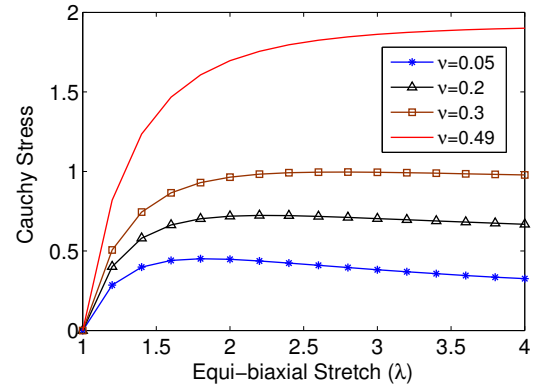
(a)



(b)



(c)



(d)

Figure 3: (a) Contours of the normalized SED function, W_w/μ_w , in $\lambda_1-\lambda_2$ plane with $\nu = 0.3$, (b) the same W_w/μ_w of (a) as a 3D surface plot, (c) the uniaxial stress-deformation behavior for various ν , and (d) the equi-biaxial stress-deformation behavior at various ν . The Cauchy stresses are normalized with respect to the shear modulus μ_w .

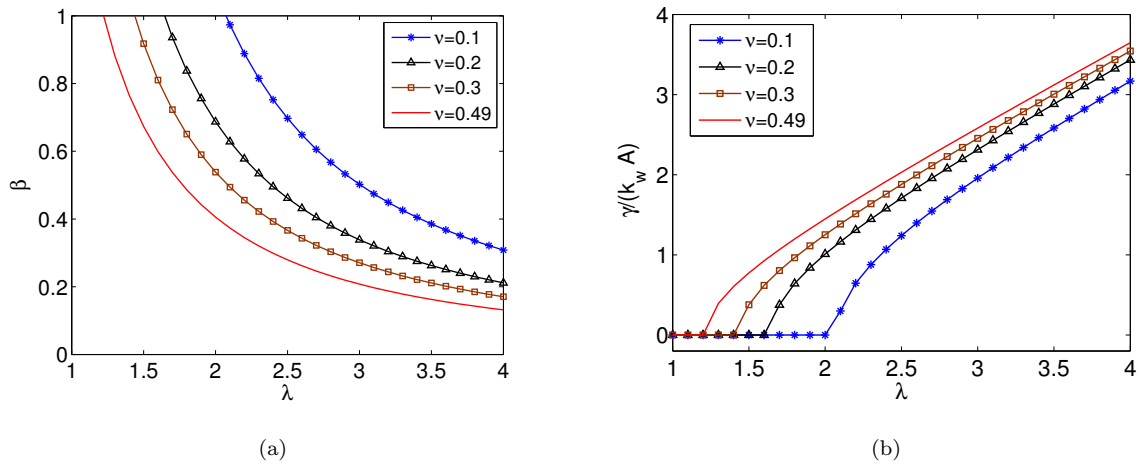


Figure 4: (a) The variation of stretch ratio $\beta = \lambda_1/\lambda_2$ with applied stretch for different ν . (b) The variation of normalized void radius $\gamma/k_w A$ with applied stretch for different ν . It is only after a critical value of applied stretch that voids begin to nucleate and eventually grow into a cavity of finite size.

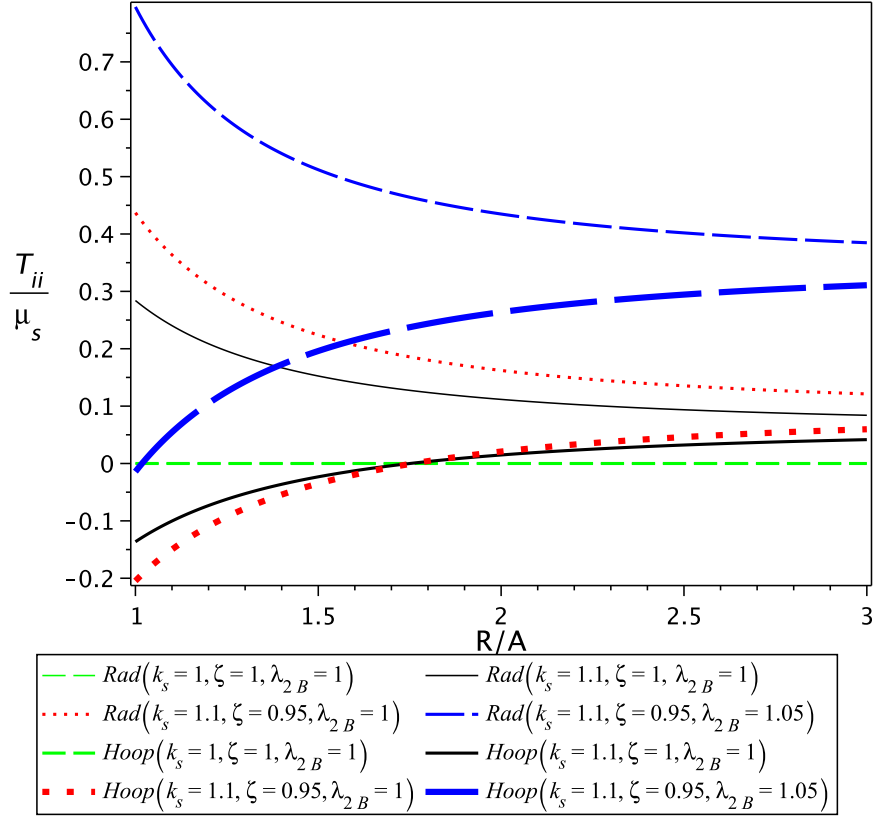


Figure 5: The normalized stresses T_{ii}/μ_s are plotted for various cases of the wound-skin arrangement, where $i = r, \theta$ for radial and circumferential stresses represented as ‘Rad’ and ‘Hoop’ in the legends. The green lines indicate stresses without any stretching, healing or incompatibility, whereas black lines show the effect of incompatibility alone, red lines show the effect of incompatibility and healing, and blue lines show the effect of incompatibility, healing, and stretching together. We fix $\alpha = B/A = 3$.

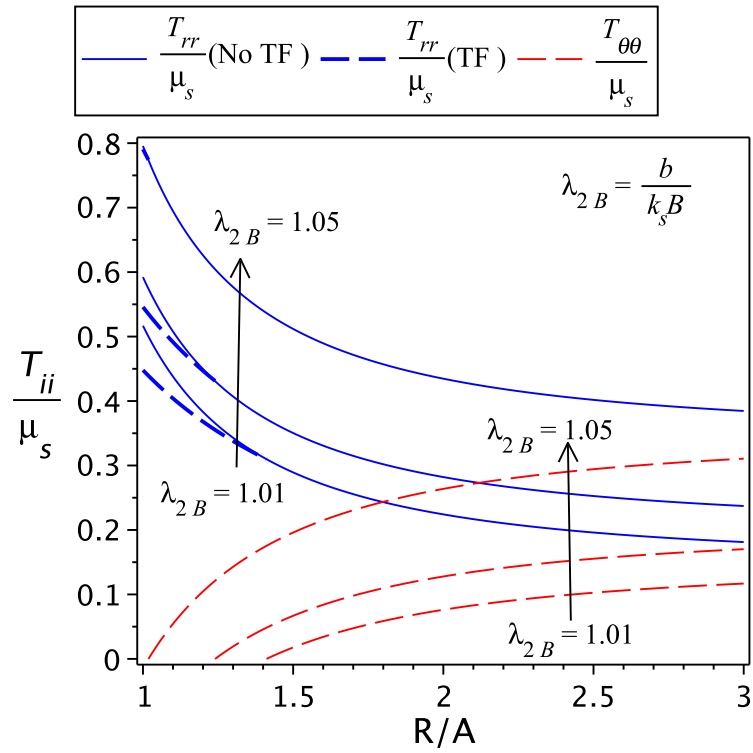


Figure 6: Normalized stresses for various applied stretches ($\lambda_{2B} = 1.01, 1.02,$ and 1.05) at the outer skin edge, with $\alpha = 3$, $\zeta = 0.95$, $k_s = 1.1$, and $\mu_w = 0.7\mu_s$. The tension field (TF) solutions are shown as dashed lines.

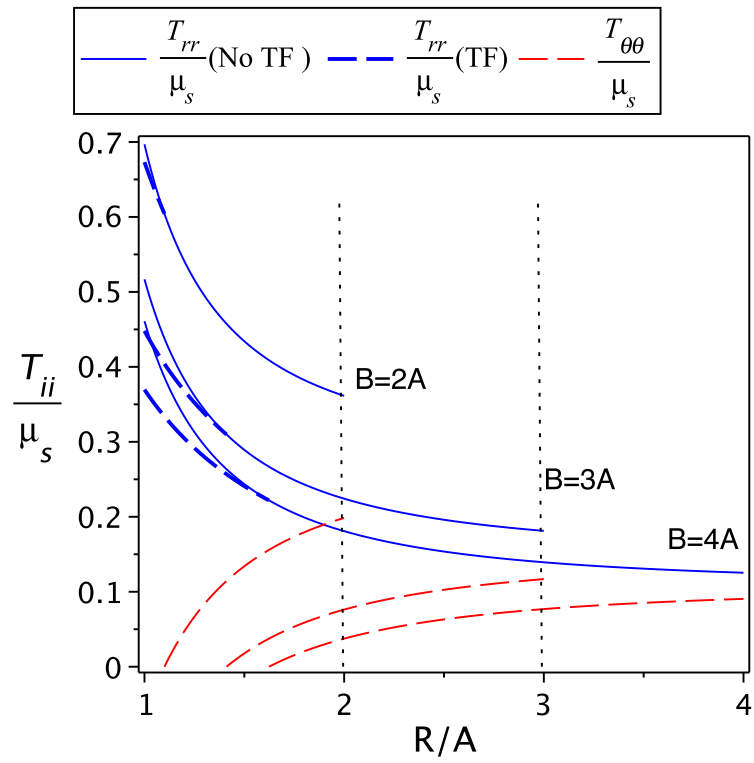


Figure 7: Normalized stresses for various positions of the applied stretch, with $\lambda_{2B} = 1.01$, $\zeta = 0.95$, $k_s = 1.1$, and $\mu_w = 0.7\mu_s$. The tension field (TF) solutions are shown as dashed lines.

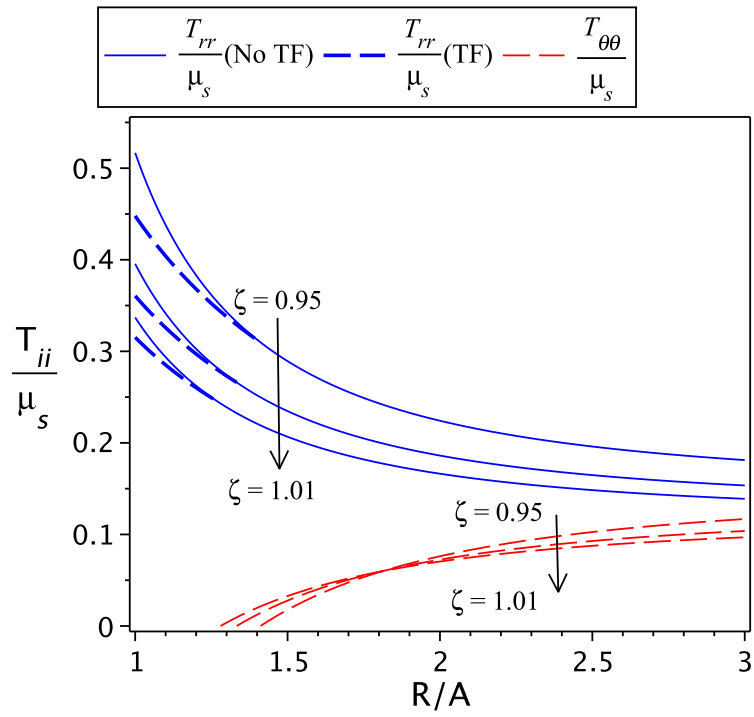


Figure 8: Normalized stresses plotted for various healing rates, where $\zeta = 0.95, 0.99$, and 1.01 represent fast healing, slow healing, and atrophy, respectively. Here, $\alpha = 3$, $\lambda_{2B} = 1.01$, $k_s = 1.1$, and $\mu_w = 0.7\mu_s$. The tension field (TF) solutions are shown as dashed lines.

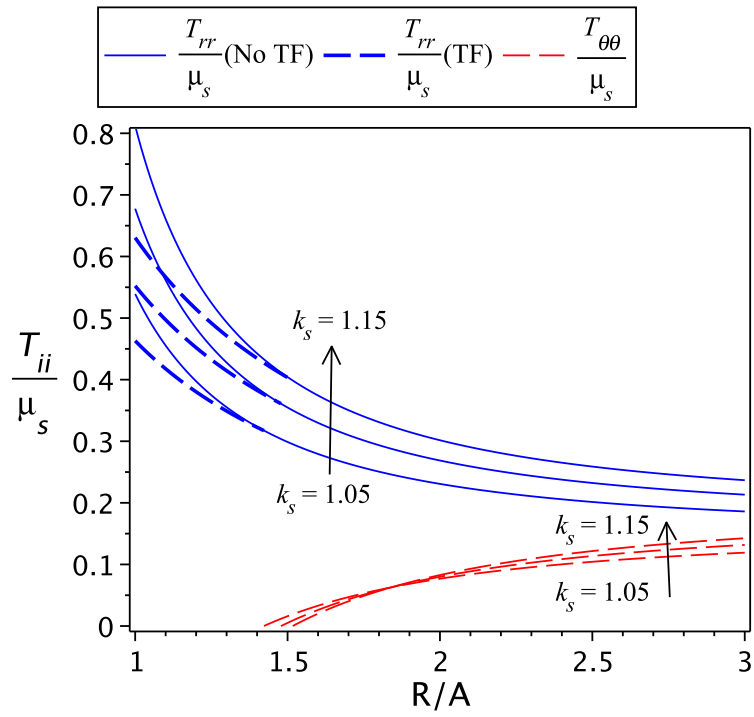


Figure 9: Normalized stresses plotted for various values of incompatibility constant ($k_s = 1.05$ (less), 1.1 (moderate), and 1.15 (high)). Here, $\alpha = 3$, $\lambda_{2B} = 1.01$, $\zeta = 0.9$, and $\mu_w = 0.7\mu_s$. The tension field (TF) solutions are shown as dashed lines.