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Role of gradients in vocal fold elastic modulus on phonation

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Abstract

New studies show that the elastic properties of the vocal folds (VFs) vary locally. In particular strong gradients exist in the distribution of elastic modulus along the length of the VF ligament, which is an important load-bearing constituent of the VF tissue. There is further evidence that changes in VF health are associated with alterations in modulus gradients. The role of VF modulus gradation on VF vibration and phonation remains unexplored. In this study the magnitude of the gradient in VF elastic modulus is varied, and sophisticated computational simulations are performed of the self-oscillation of three-dimensional VFs with realistic modeling of airflow physical properties. Results highlight that phonation frequency, characteristic modes of deformation and phase differences, glottal airflow rate, spectral-width of vocal output, and glottal jet dynamics are dependent on the magnitude of VF elastic modulus gradation. The results advance the understanding of how VF functional gradation can lead to perceptible changes in speech quality.

Keywords: phonation, vocal fold biomechanics, functional property gradation, speech quality

1. Introduction

There is evidence that the vocal fold (VF) state of health influences the spatial distribution of VF elastic properties. Kelleher et al. (2012) show that gradients in elastic modulus are smaller in cover and ligament specimens excised from subjects associated with tobacco use than specimens excised from non-smokers. Kelleher et al. (2010) show that in vacuo eigenmodes of VF tissue are dependent on gradients in elastic modulus. Zhang et al. (2007) show in vacuo eigenmodes playing an important role in the onset of flow-structure interaction (FSI). These findings lead to the hypothesis that functional gradients in VF tissue modulus influence VF dynamics during self-sustained FSI. Answering this hypothesis would contribute to understanding why several studies report perceptible differences between speech quality of smokers and non-smokers.

A very limited number of studies conduct the simulation of VF self-oscillation under conditions of three-dimensional (3D) geometry and physically reasonable air flow and VF tissue properties. These conditions

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13 impose significant computational modeling challenges. Bhattacharya and Siegmund (2014b) demonstrate
 14 the use of commercially available dedicated solvers for flow and structural domains to solve problems of
 15 VF FSI including vibration and contact, VF dehydration (Bhattacharya and Siegmund, 2014a) and surface
 16 adhesion (Bhattacharya and Siegmund, 2015). Bhattacharya and Siegmund (2014c) validated this framework
 17 against experiments on physical replicas.

18 This study aims to obtain insights into the role of gradients in VF elastic modulus on VF dynamics
 19 during phonation. FSI simulations are conducted using a partitioned approach, whereby segregated solvers
 20 for the governing equations of the solid and fluid domains exchange information after every time increment.
 21 The investigation is limited to a linear elastic isotropic description of VF tissue properties situated within
 22 a 3D model of the glottal tract. The influence of gradation is quantified by analysing VF surface dynamics
 23 during phonation.

24 **2. Method**

25 *2.1. Computational model*

26 The VF model comprises separate continuum region definitions for the glottal airflow and the pair of VFs.
 27 The FSI model describes the interaction between each VF and the airflow (Bhattacharya and Siegmund,
 28 2014b).

29 The M5 description (Scherer et al., 2001) defines the geometry of the airflow domain (figure 1a) with
 30 a rectangular x_{is} - x_{ml} - x_{ap} coordinate system aligned with inferior–superior (*is*), medial–lateral (*ml*) and
 31 anterior–posterior (*ap*) directions. The sub- and supra-glottal tracts have uniform rectangular cross-sectional
 32 dimensions (*ml*: $W = 17.4$ mm, *ap*: $L = 20.0$ mm) but unequal *is* dimensions ($T_{\text{entry}} = 10.0$ mm and
 33 $T_{\text{exit}} = 20.0$ mm respectively). The *is* dimension of the glottal region is $T = 10.7$ mm. The fluid medium
 34 has properties of air (constant density $\rho_f = 1.23$ kg/m³, dynamic viscosity $\mu = 1.79 \cdot 10^{-5}$ kg/m·s) modeled
 35 as a Newtonian fluid with fluid stress $\boldsymbol{\tau}_f$ and fluid velocity \vec{v} related by

$$\boldsymbol{\tau}_f = \mu [\nabla \vec{v} + (\nabla \vec{v})'] . \quad (1)$$

36 The continuity and momentum conservation equations are

$$0 = \oint_{\partial(V^f)} (\vec{v} - \vec{v}_g) \cdot d\vec{S} \quad (2)$$

$$\text{and } 0 = \rho_f \frac{d}{dt} \int_{V^f} \vec{v} dV + \rho_f \oint_{\partial(V^f)} \vec{v} (\vec{v} - \vec{v}_g) \cdot d\vec{S} + \oint_{\partial(V^f)} p \mathbf{I} \cdot d\vec{S} - \oint_{\partial(V^f)} \boldsymbol{\tau}_f \cdot d\vec{S}, \quad (3)$$

37 along with boundary conditions

$$p(x_{\text{is}} = -T_{\text{entry}} - T) = p_{\text{in}}(t) \quad (4)$$

$$p(x_{\text{is}} = T_{\text{exit}}) = 0, \quad (5)$$

$$\vec{v}(x_{\text{ap}} = \pm L/2) = \vec{v}_g(x_{\text{ap}} = \pm L/2) = 0, \quad (6)$$

$$\text{and } \vec{v}(x_{\text{ml}} = \pm W/2) = \vec{v}_g(x_{\text{ml}} = \pm W/2) = 0, \quad (7)$$

38 with p the fluid pressure, \vec{v}_g the discretized grid velocity and $p_{\text{in}}(t)$ the time-varying pressure at the inlet

$$p_{\text{in}}(t) \equiv p_{\text{max}} \begin{cases} (t/t_0)^2 [3 - 2(t/t_0)] & \forall t \in [0, t_0] \\ 1 & \forall t \in [t_0, \infty) \end{cases} \quad (8)$$

39 where $p_{\text{max}} = 400$ Pa and $t_0 = 0.150$ s. Zero pressure at the outlet and no-slip and no-penetration at all
 40 bounding surfaces except the inlet and outlet are enforced. Here \vec{v} represents the fluid velocity, V^f the
 41 volume of the fluid domain, $\partial(V^f)$ its bounding surface, p the fluid pressure, \mathbf{I} the second-order identity
 42 tensor, $\boldsymbol{\tau}_f$ the surface traction vector on the fluid boundary. The operator \cdot represents a tensor contraction
 43 and the operator ∇ the gradient vector. The motion of the moving-deforming glottal surface given by
 44 the grid velocity \vec{v}_g is determined by the FSI model (described later). The fluid volume is discretized using
 45 tetrahedral cells, with a minimum cell size of 0.050 mm near the glottis ensured throughout the computation.
 46 The fluid model is implemented in ANSYS/Fluent (ANSYS Fluent Release 12.0 User Guide, 2009). The
 47 solution is advanced in time following the implicit PISO (Pressure Implicit with Splitting of Operator)
 48 algorithm with neighbor and skewness correction (Issa, 1986).

49 The VF domain comprises identical and disjoint left and right solid parts (figure 1b shows the left VF).
 50 Both VFs have a depth $D = 8.40$ mm separated initially by $d_g = 0.600$ mm. VF mechanics is governed by
 51 the principle of virtual work (Zienkiewicz et al., 2005)

$$\int_{V^s} \boldsymbol{\sigma} : \delta \mathbf{D}_v \, dV = \oint_{\partial(V^s)} \boldsymbol{\tau}_s \cdot \delta \vec{u}_v \, dS - \int_{V^s} \rho_s \ddot{\vec{u}} \cdot \delta \vec{u}_v \, dV. \quad (9)$$

52 with $\boldsymbol{\sigma}$ the Cauchy stress, V^s the solid volume, $\boldsymbol{\tau}_s$ the traction applied on the boundary $\partial(V^s)$, ρ_s the
 53 uniform solid density, \vec{u} the solid displacement, $\mathbf{D} = \nabla \vec{u}$ the displacement gradient, δ a variation of the
 54 virtual variables (subscripted 'v'), operator $:$ the double-contraction of two tensors, accent-marks $\dot{}$ and $\ddot{}$
 55 respectively the first- and second-order time-derivatives and ν the Poisson's ratio. The VF constitutive
 56 behavior is isotropic linear viscoelastic with $\boldsymbol{\sigma}$ depending on the history of the deviatoric strain rate $\dot{\boldsymbol{\epsilon}}$ and
 57 bulk strain rate $\dot{\boldsymbol{\epsilon}}$

$$\boldsymbol{\sigma}(t) = \int_0^t 2G(t-t') \dot{\boldsymbol{\epsilon}} \, dt' + \mathbf{I} \int_0^t K(t-t') \dot{\boldsymbol{\epsilon}} \, dt' \quad (10)$$

58 The time dependence of the shear and bulk moduli are

$$G(t) = \frac{E}{2(1+\nu)} \left[1 - g_1 + g_1 e^{-t/\tau_1} \right], \quad (11)$$

$$\text{and } K(t) = \frac{E}{3(1-2\nu)} \left[1 - k_1 + k_1 e^{-t/\tau_1} \right] \quad (12)$$

59 The viscoelastic relaxation is modeled by shear and bulk relaxation factors $g_1 = 0.100$ and $k_1 = 0.100$
60 respectively and relaxation time-constant $\tau_1 = 0.100$ s. The elastic modulus E varies in the ap direction as

$$E(x_{ap}) = E_0 + E_1 f(x_{ap}) \quad (13)$$

61 with $E_0 = 6.00$ kPa and $E_1 f(x_{ap})$ an ap -variation around it (detailed in section 2.2). The tissue is assumed to
62 be nearly incompressible ($\nu = 0.450$) and having density $\rho_s = 1070$ kg/m³. The VF volumes are discretized
63 using first-order hexahedral elements with minimum edge length 0.110 mm near the medial surface where
64 the maximum deformation is expected. For further reference a line AB oriented in the ap direction and
65 lying on the left VF surface and a nodal location \mathbf{X}_{MC} corresponding to the mid-point of AB are defined
66 (figure 1b). Boundary conditions

$$\vec{u}(x_{ap} = \pm L/2) = 0, \quad \vec{u}(x_{ml} = \pm W/2) = 0 \quad (14)$$

67 constrain all degrees of freedom on the lateral, anterior and posterior surfaces. Displacement and traction
68 boundary conditions on the glottal surfaces (left VF: S_L , right VF: S_R , figure 1c) are determined from the FSI
69 model. The solid domain model is implemented in Abaqus/Standard (Abaqus Version 6.11 Documentation,
70 2011). The solution is integrated implicitly in time using the Hilber–Hughes–Taylor algorithm (Hilber et al.,
71 1977).

72 The FSI model defines the interaction between the glottal surfaces on the VF domain and the glottal
73 surface of the airflow domain. The FSI model applies the traction boundary condition (called the dynamic
74 boundary condition)

$$\boldsymbol{\tau}_s = (-p\mathbf{I} + \boldsymbol{\tau}_f) \cdot \hat{\mathbf{n}}. \quad (15)$$

75 ensuring equal and opposite tractions acting on the domains of glottal surfaces in both model parts. Terms
76 on the left and right sides of (15) are evaluated by interpolating between neighbouring nodes taken from
77 the VF and airflow models respectively. The FSI model computes the grid velocity of the glottal surfaces
78 in the airflow model from the kinematic boundary condition $\vec{v}_g = \dot{\vec{u}}$ where $\dot{\vec{u}}$ is evaluated on S_L and S_R .
79 The deformed glottal surface geometries in the airflow and VF models always remain coincident. The
80 dynamic and the kinematic boundary conditions are applied at intervals of 50 μ s, which equals the fixed
81 time-increment of both the flow and solid domain solvers. The FSI model is implemented in MpCCI version
82 4.1 (Joppich and Kürschner, 2006).

Table 1: Modulus distribution function $f(x_{\text{ap}})$.

x_{ap} [mm]	-10.0	-6.10	-2.40	-0.10	1.40	2.40	5.90	10.0
$f(x_{\text{ap}})$	-0.711	-0.711	0.154	3.18	0.154	0.731	-0.711	-0.711

2.2. Parametric study of gradients in elastic modulus

A spatially varying function $E(x_{\text{ap}})$ is used to define the VF elastic modulus (13). The average elastic modulus $E_0 = 6$ kPa is based on the *ap* elastic modulus of the lamina propria (comprising the cover and ligament layers) measured by Zhang et al. (2009). The *ap*-variation of $E(x_{\text{ap}})$ is modeled by $f(x_{\text{ap}}) \equiv (E_l - \bar{E}_l)/\bar{E}_l$, where the *ap*-graded elastic modulus E_l (with an average value $\bar{E}_l = 182$ kPa) was measured from an excised vocal ligament of a 60 year old male subject (Kelleher et al., 2010). From the measurements of Kelleher et al. (2010) $f(x_{\text{ap}})$ is computed (table 1) and a piecewise-linear fit is plotted in figure 2. In (13) E_1 represents the magnitude of VF modulus gradation. In this study the value of E_1 is varied parametrically. In case 1, $E_1 = 0$ kPa represents a homogeneous distribution, in case 2 an intermediate value $E_1 = 3$ kPa is used and finally in case 3, $E_1 = 6.00$ kPa ($= E_0$) creates a distribution with high heterogeneity. E_0 is identical across the three cases.

2.3. Modal analysis of in vacuo and FSI dynamics

Modal analysis of in vacuo VF dynamics is frequently used to characterize VF oscillatory behavior. It suffices to analyze only the left VF since both VFs possess identical geometry and material properties. In this analysis the boundary conditions constraining all degrees of freedom on the lateral, anterior and posterior VF surfaces remain identical to those in the FSI analysis (14), but the surface S_L is stress-free since fluid excitation is absent. Viscoelasticity of the VF tissue is ignored in this analysis. Modal analysis gives in vacuo eigenfrequencies (also known as natural frequencies) and eigenmodes.

On the other hand, VF motion in FSI is controlled by fluid loading. A well-known method for analyzing dynamic data, such as from FSI, is proper orthogonal decomposition (POD). POD analysis is conducted for only the left glottal surface displacements, thereby neglecting left-right asymmetries in VF motion. Left VF displacement data is used to populate a matrix, with rows corresponding to 3060 glottal surface nodes and columns corresponding to over 30 distinct time instants within a given vibration cycle. Singular value decomposition of the matrix yields for every k -th FSI mode an eigenvalue $\alpha^{(k)}$ and corresponding modes shapes $\vec{h}^{(k)}(\vec{x})$ (spatial) and $g^{(k)}(t)$ -s (temporal). FSI modes are numbered by the descending order of their eigenvalues. Only the first three FSI modes were analyzed. The spatio-temporal variation of VF displacement is approximated as

$$\vec{u}(\vec{x}, t) \simeq \sum_{k=1}^3 \alpha^{(k)} \vec{h}^{(k)}(\vec{x}) g^{(k)}(t). \quad (16)$$

Table 2: Influence of elastic gradients on FSI frequency F_0 and medial–lateral displacement at mid-coronal location $u_{ml}(\mathbf{X}_{MC})$ averaged over multiple vibration cycles (with standard deviations SD), and on the first three FSI eigenvalues $\alpha^{(k)}$ ($k = 1 \dots 3$) obtained from POD analysis of vibration cycles specified by start and end time instants t_{start} and t_{end} respectively.

Case	1	2	3
F_0 [Hz]	108 (SD 4.18 %)	109 (SD 4.95 %)	153 (SD 3.48 %)
$u_{ml}(\mathbf{X}_{MC})$ [mm]	−0.212	−0.198	−0.201
t_{start} [s]	0.2538	0.1968	0.2838
t_{end} [s]	0.2639	0.2054	0.2900
$\alpha^{(1)}$	0.349	0.210	0.212
$\alpha^{(2)}$	0.106	0.027	0.018
$\alpha^{(3)}$	0.050	0.019	0.015

Parseval’s identity states that the square of $\alpha^{(k)}$ divided by the sum of squares of all $\alpha^{(k)}$ -s is the fraction of the total spatio-temporal variation of displacement explained by the k -th FSI mode.

3. Results

3.1. *in vacuo* eigenmodes

The first three *in vacuo* eigenfrequencies (natural frequencies) are: in case 1, 48.2 Hz, 65.6 Hz and 79.1 Hz; in case 2, 53.2 Hz, 64.6 Hz and 78.7 Hz; and in case 3, 47.7 Hz, 50.3 Hz and 61.0 Hz. Across cases 1–3 the first eigenfrequency varies only slightly from 47.7 to 53.2 (10.9 % difference). This correspondence is maintained between cases 1 and 2 even for the next two eigenfrequencies (difference not exceeding 1.54 %) which are somewhat higher than the respective eigenfrequencies of case 3.

3.2. Local glottal surface motion characteristics in FSI

The *ml* displacement of location \mathbf{X}_{MC} with time ($u_{ml}(\mathbf{X}_{MC})$) is considered to be representative of the motion of the left VF. Table 2 compares overall vibration characteristics for multiple cycles corresponding to cases 1–3. Average $u_{ml}(\mathbf{X}_{MC})$ does not differ significantly across the cases (SD 3.62 %). This is expected since the stiffness of the VFs is on average identical across the cases. Vibration cycles are defined to start and end at consecutive peaks of $u_{ml}(\mathbf{X}_{MC})$ in time. The phonation frequency F_0 was obtained as

$$F_0 = \frac{1}{t_{cycle}}, \quad (17)$$

where t_{cycle} separates two consecutive peaks of $u_{ml}(\mathbf{X}_{MC})$. The average F_0 is significantly different in case 3. In each case F_0 has no apparent correspondence with the first three eigenfrequencies noted earlier. For

127 case 1, the net mass flow rate through the outlet averaged 0.676 g/s, and oscillated with a frequency of
128 108 Hz (similar to F_0).

129 Next, consider one particular cycle of vibration in each of the cases 1–3. Table 2 gives corresponding to
130 each cycle, the cycle start and end times t_{start} and t_{end} respectively. Within each cycle, four representative
131 time instants are identified. Two of these instants correspond to times when $u_{\text{ml}}(\mathbf{X}_{MC})$ attains the minimum
132 and maximum value over the vibration cycle. Since negative values of $u_{\text{ml}}(\mathbf{X}_{MC})$ correspond to an opening
133 of the glottal gap, the above instants can respectively be taken to represent the open and closed states of
134 the VF. At the other two instants, $u_{\text{ml}}(\mathbf{X}_{MC})$ has intermediate values. One instant is in the closing phase
135 while the other is in the opening phase.

136 Figure 3a,b shows the variation of *is* and *ml* displacement components along the *ap* line *AB* at the four
137 representative instants for case 1. Plots for case 2 and 3 are given in figures 3c,d and 3e,f. The location
138 \mathbf{X}_{MC} corresponds to $x_{\text{ap}} = 0$. Closed and open instants and the intermediate instants are distinguished in
139 these figures. Net mass flow rates through the outlet in dependence of time are shown.

140 3.3. POD analysis

141 Table 2 gives the first 3 FSI eigenvalues from a POD analysis. In each case POD analysis is performed
142 on the displacements corresponding to the vibration cycle selected above. From Parseval’s identity the first
143 three FSI modes were found to explain at least 99.8 % of the spatio-temporal variation of the displacement
144 during the cycle. Excluding the variation explained by FSI mode 1, the remainder is explained up to
145 99.0 %, 93.7 % and 90.6 % respectively in cases 1, 2 and 3 by FSI modes 2 and 3 combined. Figures 4 and 5
146 respectively show the spatial and temporal variation of the FSI-modes. In order to facilitate comparison,
147 both $\vec{h}^{(k)}$ and $g^{(k)}$ are normalized to lie within $[-1, 1]$. The correlation between FSI modes and in vacuo
148 eigenmodes is given in table 3.

149 3.4. Airflow

150 Figure 6a shows the variation of mass-flow rate with time for the selected vibration cycle in each case. In
151 order to facilitate comparison both mass flow rate and time ranges are normalized to lie in the unit interval
152 (figure 6b).

153 Figure 7 shows, at closed and open states, the region of the glottal jet possessing a high vorticity
154 magnitude ($|\omega| = |\nabla \times \vec{v}| \geq 10^4 \text{ s}^{-1}$). The jet region boundary is colored by contours of velocity magnitude
155 $|\vec{v}|$.

156 4. Discussion

157 The in vacuo frequencies for case 1 (no elastic gradients) compare excellently with those determined
158 by Zhang (2011) for a nearly identical VF model. The small differences are attributed to $E_0 = 4 \text{ kPa}$ in Zhang

Table 3: Square of correlation R^2 between FSI-modes and in vacuo eigenmodes. A dash indicates a value smaller than 0.001.

		in vacuo mode 1	in vacuo mode 2	in vacuo mode 3
Case 1	FSI mode 1	0.945	–	0.132
	FSI mode 2	0.006	0.003	0.015
	FSI mode 3	0.092	0.013	0.064
Case 2	FSI mode 1	0.920	–	0.144
	FSI mode 2	0.008	0.042	0.031
	FSI mode 3	0.032	0.031	0.034
Case 3	FSI mode 1	0.872	0.014	0.136
	FSI mode 2	–	0.018	0.017
	FSI mode 3	0.001	0.162	–

159 (2011) as against 6 kPa in this paper. Since E_0 is fixed for cases 1–3, the first in vacuo eigenfrequency and the
160 average displacement (during FSI) are independent of elastic gradients (table 2). However, elastic gradients
161 strongly influence the phonation frequency F_0 in FSI (table 2). This follows the argument of Zhang et al.
162 (2007) that F_0 is dependent on a flow-induced stiffness. The complexity of the flow–structure interaction
163 perhaps justifies the non-linear increase in F_0 from case 1 to case 3.

164 Average F_0 (table 2), ml displacement (table 2) and net mass-flow rate through the glottis were within the
165 range of observations in experimental studies conducted with similar VF and glottal tract geometry (Alipour
166 and Scherer, 1995; van den Berg et al., 1957; Cranen and Boves, 1985; Erath and Plesniak, 2006, 2010; George
167 et al., 2008; Morris and Brown Jr., 1996; Scherer et al., 2001; Thomson et al., 2005; Titze, 2006; Triep et al.,
168 2005; Triep and Brücker, 2010; Zhang et al., 2006).

169 Considering the variation of is displacement along AB at different instants (figures 3a,c,e), the average
170 deformed shape goes from an inverted-U shape in case 1 to an M shape in case 3. The increase in gradient
171 of elastic modulus from case 1 to 3 is associated with a decrease in stiffness near the anterior and posterior
172 ends and an increase in stiffness in the mid-membranous region. This causes more bulging near the ends
173 than in the middle section. A similar behavior was predicted in Kelleher et al. (2010) where the analysis
174 was based on a study of in vacuo mode shapes. The time-dependence of the deformed shape is obtained
175 uniquely in FSI and is analyzed further below.

176 In the insets of figure 3 the mass-flow rate is plotted throughout the vibration cycle from which the
177 displacement graphs in figure 3 are obtained. Though the frequency of oscillation of mass-flow rate and of
178 $u_{ml}(\mathbf{X}_{MC})$ were very close to each other, there exists a noticeable phase difference between the two. The
179 phase difference can be attributed to the 3D nature of VF vibration, i.e. the net mass flow rate depends

180 on u_{ml} at more than one ap location. In cases 1 and 2, the flow rate leads the displacement, whereas the
 181 situation is reversed in case 3. In case 3, u_{ml} plotted along line AB at different instants within a vibration
 182 cycle (figure 3f) suggest that while the mid-membranous region is in closing phase, a slightly off-center
 183 location could be in the opening phase. This can substantially alter the dynamics of the glottal jet, and in
 184 particular the phase difference.

185 Focusing on FSI mode 1, table 2 shows that in each case this mode has the highest eigenvalue, implying
 186 most of the deformation of the VF is captured by FSI mode 1. Figure 5 shows that the temporal variation
 187 of FSI mode 1 is negligible, with $g^{(1)}(t)$ possessing values close to -1 . Hence, the FSI mode 1 describes
 188 the non-oscillating part of deformation. The FSI mode 1 shape (figure 4), after multiplying with the -1
 189 factor, captures well the average upward bulge of the VF around which further oscillation takes place. This
 190 also agrees with the average is displacement (figures 3a,c,e). In each case, the variation in FSI mode 1 is
 191 explained most by in vacuo modes 1 and 3, in order.

192 Figure 4 shows FSI mode 2 characterized by oscillatory bulging out and shrinking in of the glottal
 193 surface, primarily in the ml direction. Going from the homogeneous VF (case 1) to the highly graded VF
 194 (case 3), the peak bulge location shifts from a mid-coronal position to two increasingly separated anterior
 195 and posterior positions. This agrees with the variation of u_{ml} along AB (figures 3b,d,f), and is explained
 196 directly by the variation of $E(x_{ap})$ across the three cases.

197 FSI mode 3 causes an upward bulge varying in the ml direction. While the lateral regions dip down, the
 198 medial surface lifts up, and vice versa. As in the case of other FSI mode shapes, the coronal plane at which
 199 the downward dip and upward lift are maximum moves from a mid-coronal location in case 1 and splits into
 200 two anterior and posterior located coronal planes in case 3.

201 Figure 5 indicates that in each case $g^{(k)}(t)$ is on average zero for FSI modes 2 and 3 implying a purely
 202 oscillatory behavior. The frequency of oscillation with time equals F_0 . Interestingly $g^{(2)}(t)$ leads $g^{(3)}(t)$ by
 203 90° in cases 1 and 2, but then lags by the same phase in case 3.

204 After excluding the part explained by FSI mode 1, a decreasing fraction of VF dynamics is explained by
 205 FSI modes 2 and 3 combined as one goes from case 1 to case 3. This suggests that increase in VF elastic
 206 gradients leads to an increase of the number of distinct FSI modes recruited in VF oscillation, and thereby
 207 to an increase in spectral width of the oscillation signal.

208 For all 3 cases, the variation in FSI modes 2 and 3 explained by the first three in vacuo eigenmodes is
 209 below 17 % (table 3). This indicates that higher order in vacuo modes make significant contributions to FSI
 210 modes 2 and 3. For FSI mode 3 in case 1, the first and third in vacuo eigenmode make higher contributions
 211 than the second in vacuo eigenmode. These relative contributions to FSI mode 3 tend to become more
 212 uniform in case 2. Finally in case 3 a complete reversal is achieved: the in vacuo mode 2 makes a higher
 213 contribution to FSI mode 3 than the in vacuo modes 1 and 3.

214 Comparing the in-cycle mass flow rate (figure 6a) in cases 1 and 2, it is clear that the fluctuation amplitude

215 (or contrast) of the flow is higher for case 2 with higher gradient. This emphasises the importance of a more
216 trapezoidal opening in case 2 as mentioned before. This confirms a similar prediction made in Kelleher et al.
217 (2010) following a linear eigenmode analysis.

218 The normalized waveforms in figure 6b suggest that a higher gradient of elastic modulus can induce
219 differences in the spectrum of harmonic excitation provided by the airflow output. This agrees with the
220 previous observation that increase in gradients of elastic modulus increase the spectral width of VF dynamics
221 in FSI. Flow rates in cases 1 and 2 have a noticeably different spectral-width than the flow rate in case 3.
222 Since collision between the VFs was not considered, the LF pattern (Fant and Liljencrants, 1985) is not
223 apparent. Rather the airflow curves are similar to those obtained for breathy voice (Hillenbrand and Houde,
224 1996).

225 Figure 7 demonstrates the effect of VF vibration shapes on the glottal jet shape. The jet structure
226 changes significantly in the closed state. Especially in cases 2 and 3, the *ap* asymmetry in displacements
227 caused by the gradients in elastic modulus is reflected in the jet passing through the glottis. In case 1 the
228 flow is reduced in the mid-membranous region, resulting the jet to split in the closed state. However in
229 cases 2 and 3, the jet is preferentially aligned to one side along the *ap* direction. The Coanda effect is well
230 documented in VF flows, and is known to cause the jet to move in the *ml* direction. In this study, the
231 Coanda effect was observed though it is not shown here. The *ap* asymmetry of the jet (expected to be
232 substantial when gradients in elastic modulus exist) along with the Coanda effect will cause the glottal jet
233 to have an exceedingly high 3D character.

234 5. Conclusion

235 This study was conducted to investigate the role of gradients in elastic modulus on VF dynamics and
236 airflow. Elastic modulus variation along the *ap* direction of the vocal ligament, obtained by careful mea-
237 surements based on digital image correlation, were used to inform a full 3D VF tissue model. Specifically,
238 elastic modulus within the VF tissue possessed a variation similar to that measured in the ligament, but the
239 average modulus was based on a study that measured the average modulus of the lamina propria. An FSI
240 investigation was conducted with three models with successively increasing gradients in elastic modulus. A
241 study on the effect of elastic modulus gradients has never been attempted before, and hence the present
242 results provide novel insights.

243 An important observation is that changes in gradients of elastic modulus alter the phonation frequency
244 in a highly non-linear fashion. An increase in the gradient can lead to the following changes in VF dynamics,

- 245 • a more trapezoidal deformed shape of the VF in the *is* direction,
- 246 • higher amplitude of vibration at off-mid-membranous locations,

247 • increased *ap* asymmetry in *is* and *ml* displacements, and

248 • *ap* mucosal waves (see supplementary material).

249 POD analysis of VF dynamics suggests that elastic gradients influence

250 • the number of distinct FSI modes recruited in vibration

251 • the phase relationship between the modes

252 • the correspondence of FSI modes with in vacuo eigenmodes

253 These changes are concomitant with following alterations in glottal flow

254 • higher contrast of glottal airflow output,

255 • increase in spectral-width of glottal airflow output, and

256 • strong *ap* asymmetry of the glottal jet.

257 Phonation frequency, contrast and spectral-width of glottal airflow output and glottal jet dynamics influence
258 speech quality significantly (Rotherberg, 1981; Titze and Story, 1997). The present study demonstrates a
259 strong link between gradients in elastic modulus and speech quality.

260 The main limitations of this study arise out of modeling simplifications. The distribution of elastic
261 modulus within the length of the ligament was used as a model for the distribution within the entire VF
262 tissue. Kelleher et al. (2012) found this distribution itself changes in cover and ligament layers. Future
263 studies conducted with layered VF tissue models could provide further insights. Determining the sensitivity
264 to functional gradation in the presence of anisotropy could be interesting. The present study did not consider
265 collision between the vocal folds, but this is no principal limitation. Yet, gradients in elastic modulus can
266 be expected to play a significant role in modifying stress distributions within the VF tissue under collision
267 conditions.

268 **Conflict of interest statement**

269 The authors of this manuscript have no conflicts of interest related to the content of the study.

270 **Acknowledgement**

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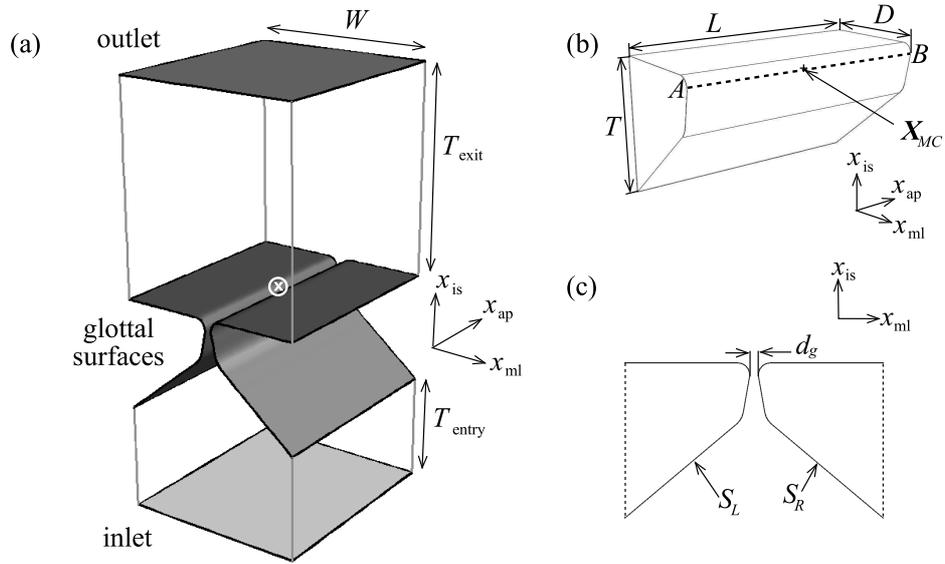


Figure 1: (a) Geometry of the glottal airflow domain: the inlet, outlet and glottal surfaces are shaded, the coordinate origin (at the intersection of the mid-coronal plane, the mid-sagittal plane and the VF superior surface) is \otimes ; (b) geometry of the left half of the solid VF model: line AB and point \mathbf{X}_{MC} are reference regions at which the VF motion is characterized in this paper; (c) mid-coronal section showing both pairs of VFs and rigid planes: coordinate axes are offset from the origin for clarity

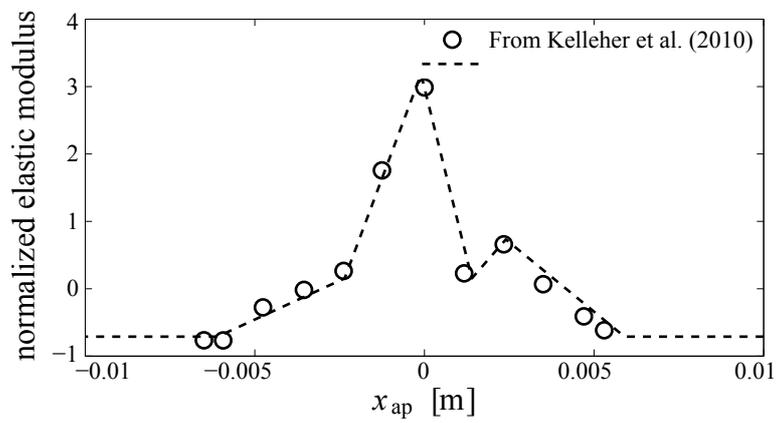


Figure 2: Piecewise linear fit to normalized elastic modulus $f(x_{ap})$ in dependence of anterior–posterior location x_{ap} . Circles correspond to values in table 1.

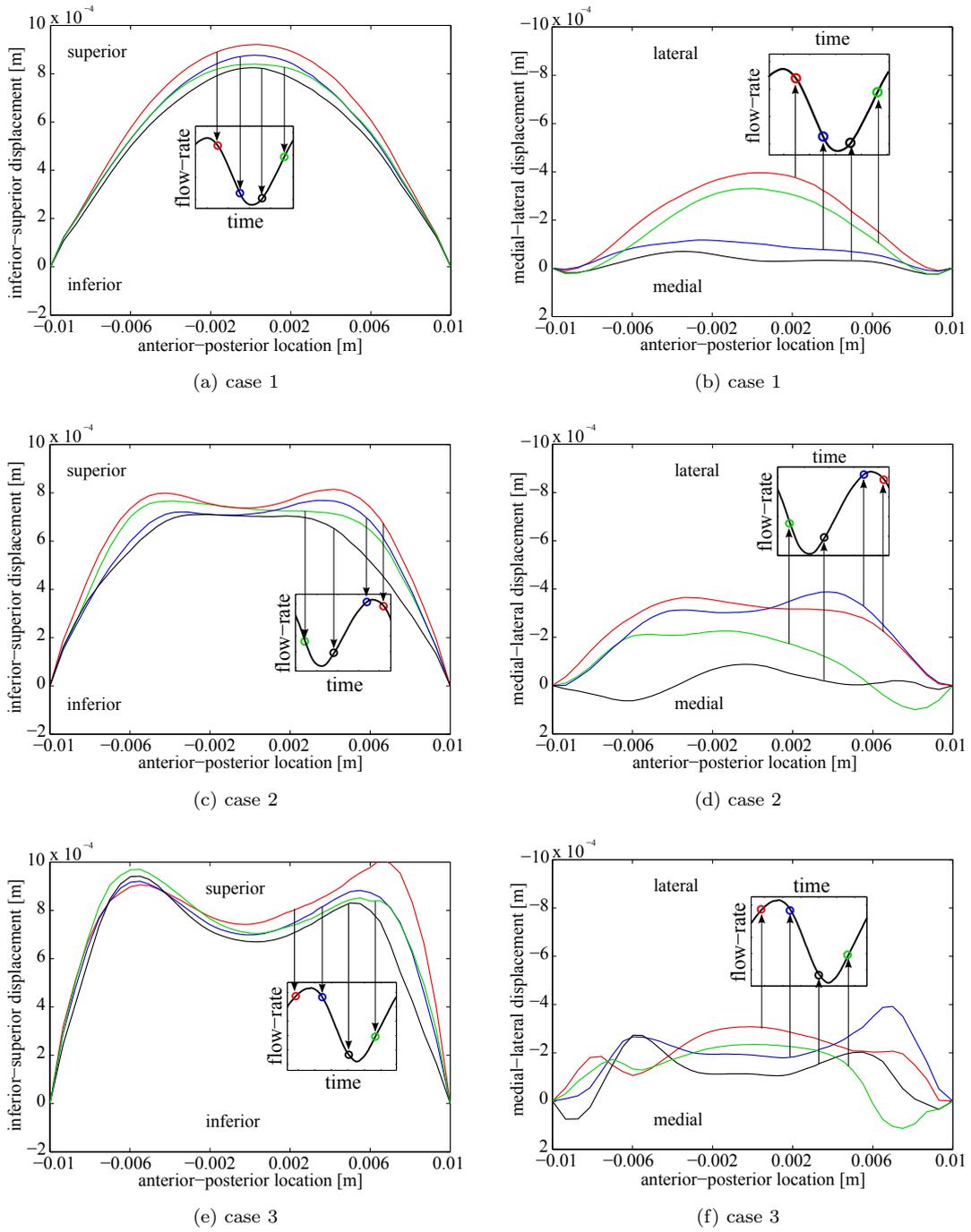


Figure 3: Displacements in inferior-superior (a,c,e) and medial-lateral (b,d,f) directions plotted along line AB at 4 instants of a vibration cycle. A more open VF configuration corresponds to a more positive inferior-superior displacement and a more negative medial-lateral displacement.

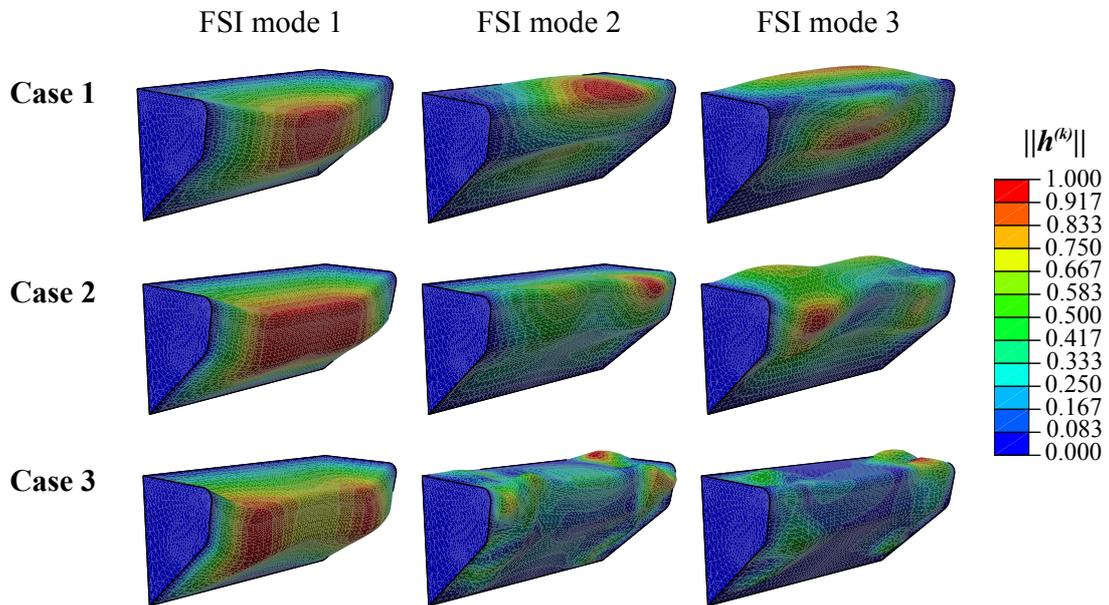


Figure 4: Spatial variation of $\vec{h}^{(k)}$ for FSI-modes $k = 1 \dots 3$. The variation is normalized by the maximum magnitude of $\vec{h}^{(k)}$.

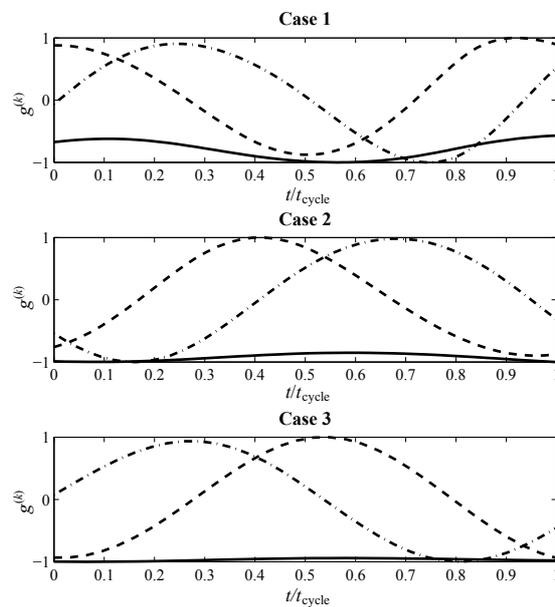


Figure 5: Temporal variation of $g^{(k)}$ for FSI-modes $k = 1 \dots 3$: $-$, $k = 1$; $- -$, $k = 2$; $- \cdot -$, $k = 3$. The scalar function $g^{(k)}(t)$ is normalized with respect to its maximum occurring magnitude.

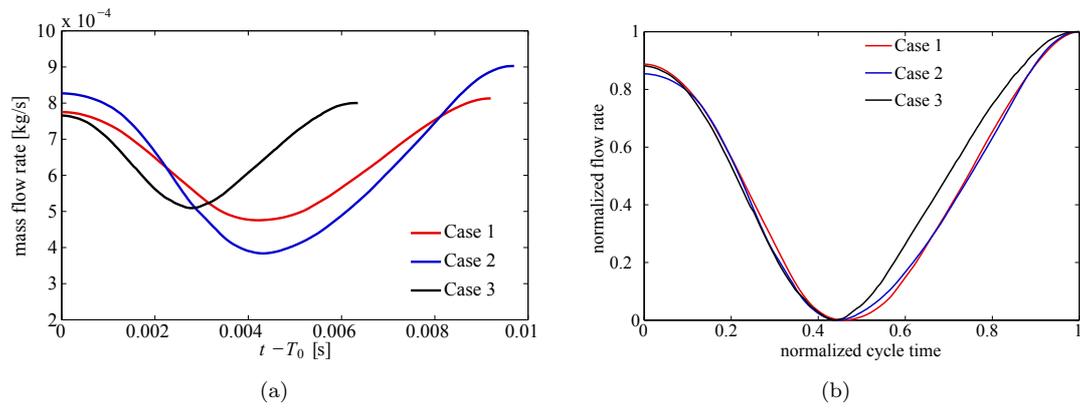


Figure 6: (a) Mass flow rate in dependence of time over one vibration cycle. Time origin is at the start time t_{start} of the corresponding cycle. (b) Curves in (a) normalized such that flow rate and cycle time are in $[0, 1]$.

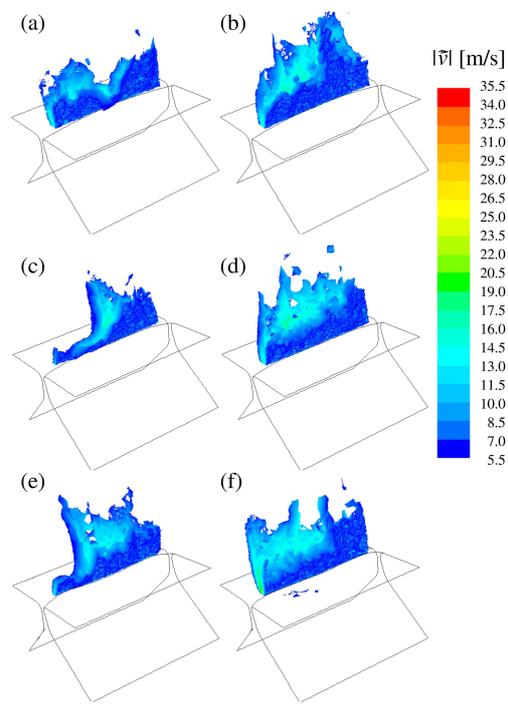


Figure 7: Velocity magnitude contours in the glottal jet: (a,b) case 1; (c,d) case 2; (e,f) case 3; (a,c,e) closed states; (b,d,f) open states.