



Imaging of auto-oscillating vocal folds replicas with left–right level difference due to angular asymmetry[☆]

Anne Bouvet^a, Isao Tokuda^b, Xavier Pelorson^a, Annemie Van Hirtum^{a,*}

^a LEGI, UMR CNRS 5519, Grenoble Alpes University, France

^b Dep. Mech. Eng., Ritsumeikan Univ., Nojihigashi, Kusatsu, Shiga 525-8577, Japan

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ABSTRACT

Vertical level difference due to vertical angular asymmetry (up to 20°) between the normal (right) and tilted (left) vocal fold (VF) is experimentally assessed in the framework of unilateral vocal fold paralysis. The upstream pressure and high-speed vibration imaging for different camera view angles are assessed during steady-state auto-oscillation of three different mechanical VF replicas. A global image-based correlation allows to assess the oscillation frequency independently from the camera view angle. Resulting oscillation frequencies matches the ones obtained from aerodynamic data. Mean vibration cycles obtained from videokymographic line-scans at different positions along the posterior–anterior direction are analysed. Left–right VF displacement phase difference, axis shift and left–right mucosal wave phase difference are identified as essential parameters quantifying angular asymmetry. The influence of the degree of angular asymmetry, scan position and replica is discussed.

1. Introduction

Unilateral vocal fold paralysis (UVFP) is a common vocal fold (VF) pathology, reported for instance to affect about 0.5% of the population in England [1–3]. In about 83% of UVFP cases dysphonia is found to reduce the quality and performance of the voice [1–3]. Clinical UVFP examinations often reveal an air escape due to left–right VF asymmetries with respect to VF's shape, tension and positioning [1,4,5]. Despite the continuous advancement of measurement techniques [6–9] quantitative accurate *in-vivo* clinical data assessment on human speakers remains tedious, which hampers a systematic assessment of the influence of these asymmetries on voice properties. As a result, a consensus concerning the definition, diagnosis and hence treatment of UVFP remains yet to be achieved [3]. From this perspective, physical studies using deformable vocal folds replicas can provide new insights pertinent to UVFP as they facilitate measurement of variables of interest as well as a systematic variation of the degree of UVFP.

Recently, a physical study using mechanical VF replicas (Fig. 1) was presented in order to experimentally study the effect of air leakage solely due to vertical left–right angular asymmetry on features commonly associated with voice quality [10]. Concretely, the right VF was kept in place (the normal one) whereas the left VF (the paralysed one) is tilted with asymmetry angle α in its sagittal plane as illustrated in

Fig. 2. Experiments were performed using three different (M5, MRI and EPI) deformable multi-layer silicone VF replicas [11–15] depicted in Fig. 1 [10,16]. Increasing the asymmetry angle α (up to 25°) increases the glottal leakage area (red triangle in Fig. 2). The shape and tension of each VF are unaltered by the imposed asymmetry angle α . It followed that the degree of contact between the medial surfaces of the left and right VF decreases and eventually ceases with α [10,16].

For all three replicas, it was found [10,16] that when α and hence glottal leakage increases, the oscillation frequency f_0 reduces monotonically with at least 10% while both VFs remain in partial contact and even more so when contact between both VFs ceases. This finding offers an interesting new perspective on frequency changes related to UVFP. Indeed, whereas clinical studies generally attribute its symptomatic low frequency voice solely to decreased tension in the paralysed vocal fold [1,17], this physical study [10] shows that frequency reduction might be partly correlated to increased air leakage due to angular asymmetry. In addition, it was found that when full contact is lost so that air leakage occurs, the oscillation quality reduces as the signal-to-noise ratio decreases, the total harmonic distortion rate increases, oscillation threshold pressures at onset and offset increase and higher harmonics become apparent. These observations bear similarities to clinical UVFP

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* Corresponding author.

E-mail address: annemie.vanhirtum@univ-grenoble-alpes.fr (A. Van Hirtum).

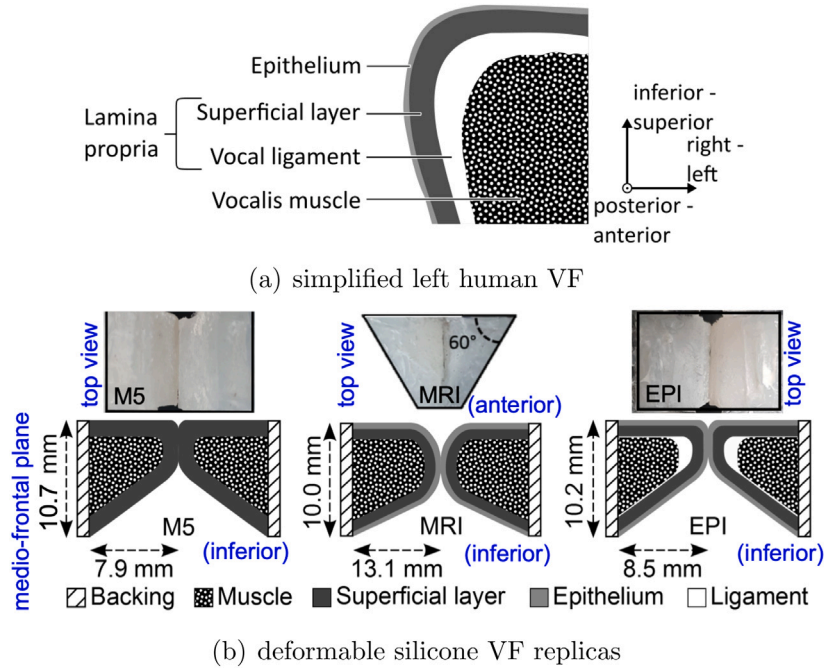


Fig. 1. Multi-layered VF structure: (a) left human VF in the medio-frontal plane based on [1], (b) deformable silicone VF replicas M5, MRI and EPI [10] for $\alpha = 0^\circ$.

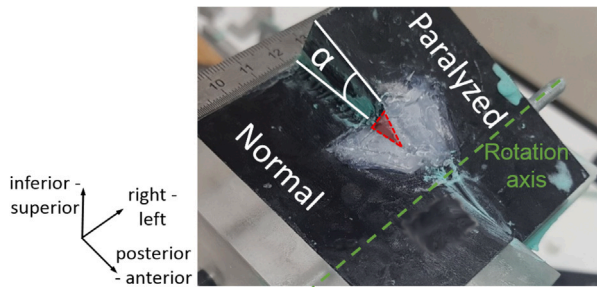


Fig. 2. Illustration of imposed asymmetry angle α , resulting glottal gap associated with air leakage (red triangle) and fixed rotation axis used for tilting of the left VF in case of the MRI VF replica with $\alpha = 20^\circ$ [10].

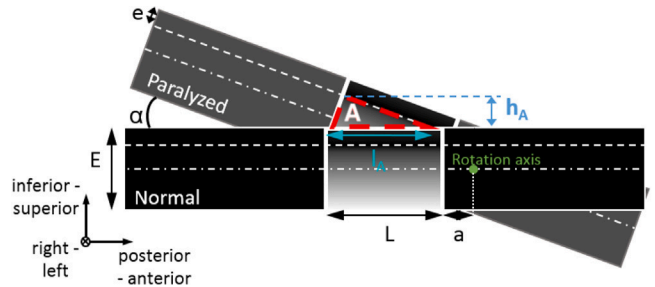


Fig. 3. Schematic sagittal side view of glottal replica and dimensions: imposed asymmetry angle α , glottal gap (red triangle) characteristics A , l_A , h_A and VF parameters E , e , L and constant rotation axis position $a = 4.5$ mm [10].

voice descriptions of dysphonia (increased harmonic distortion rate and reduced frequency), breathy voice (reduced signal-to-noise ratio), vocal fatigue and reduced maximum phonation time (increased onset pressure threshold and thus oral airflow during phonation) [1,17,18]. As the loss of full glottal contact is seen to trigger these feature changes, experimental observations support voice therapy strategies aiming to remedy glottal closure [18].

Given the described similarities, it is aimed to further investigate the effect of imposing angular asymmetry α on auto-oscillating VF replicas. As in [10,16] different deformable replicas (M5, MRI, EPI) are used in order to assess the generality of observed tendencies. Features discussed in [10,16] were obtained from analysing the subglottal pressure which is an aerodynamic quantity. One might argue that this pressure is difficult to measure directly on human speakers and that aerodynamic observations do not provide a direct measurement of the ongoing vibration. Therefore, the aim of this work is to consider the effect of the imposed angular asymmetry directly on the vibration of auto-oscillating VF replicas using high-speed imaging and to quantify clinically relevant markers. Global and local vibration features are assessed considering instantaneous images (global) or videokymographic (VK) [19] line-scans (local) and the influence of camera view angle (global) and image scan position (local) on quantified features is considered. Local VK features are sought to inform on VF asymmetry.

2. Glottal replicas

2.1. Deformable silicone VF replicas

Oscillation is studied for three different deformable multi-layered moulded silicone VF replicas [10,16] – M5, MRI and EPI shown in Fig. 1(b) – The replicas differ in geometry and composition as respectively two (M5), three (MRI) or four (EPI) moulding layers are added to a backing layer attaching it to a rigid support, i.e. the region outside of the top view frames shown in Fig. 1(b). The M5 VF replica is a two-layer (muscle and superficial layer) reference model following the so called M5 geometrical VF model [20]. The MRI VF replica has a more realistic geometry derived from magnetic resonance imaging data of a human VF [11,14]. It has a three-layer structure by adding a third thin and stiff surface layer representing the epithelium to the two-layer structure of the M5 VF replica. The EPI VF replica is obtained by inserting an extremely soft deep layer between the muscle and the superficial layer of the three-layer structure used for the MRI VF replica [13]. The EPI VF replica cast is inspired by the geometrical M5 VF model so that its geometry is a scaled version of the M5 VF replica. A detailed overview of layer thicknesses and Young modulus for each VF replica is provided in [10,16].

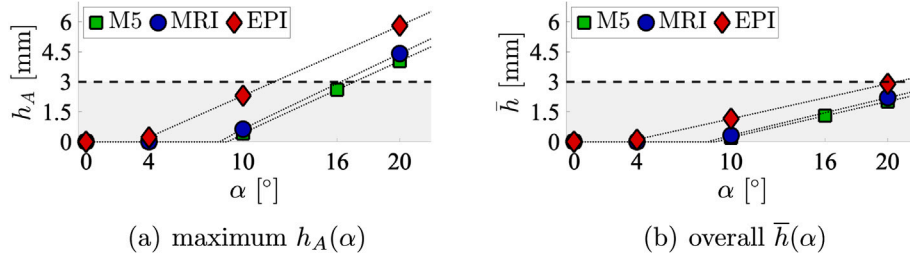


Fig. 4. Assessed vertical level differences for different replicas M5 (□), MRI (○) and EPI (◇) and range for human UVFP subjects [5] (grey-shaded region up to 3 mm): (a) maximum $h_A(\alpha)$, (b) overall $\bar{h}(\alpha)$.

Table 1

Critical angles ($\alpha_{I,II}$, $\alpha_{II,III}$) and contact regime (I, II, III) of α .

| Replica | $\alpha_{I,II}$ [°] | $\alpha_{II,III}$ [°] | I | II | III |
|---------|---------------------|-----------------------|--------|---------------|-----|
| M5 | 8.1 | 29.2 | 0°, 4° | 10°, 16°, 20° | – |
| MRI | 7.8 | 28.8 | 0°, 4° | 10°, 20° | – |
| EPI | 3.0 | 14.0 | 0° | 4°, 10° | 20° |

The initial glottal area, i.e. spacing between VFs without airflow, with left–right symmetry ($\alpha = 0^\circ$) yields less than 10 mm² for each VF replica. Mechanical resonances were identified from frequency response functions. For each replica, a first f_1^M (quality factor ≈ 11) and second f_2^M (quality factor ≈ 29) mechanical resonance peak frequency was observed at 142 ± 5 Hz and 263 ± 5 Hz, respectively.

2.2. Glottal replicas with vertical angular asymmetry angle α

Each deformable silicone VF replica is mounted so that its left VF can be tilted with angle α in the sagittal plane around a fixed rotation axis in the right–left direction. This results in a vertical level difference between the left and right VF parameterised by the imposed angular asymmetry angle α as illustrated in Fig. 2 and in Fig. 3. As the rigid support containing the VF is rotated as well, the elasticity and the shape of each VF is not altered.

As α increases, a triangular glottal air leakage gap (red triangle in Fig. 2) emerges near the posterior VF edge, which is characterised by its area A , triangle base l_A and triangle height h_A indicated in Fig. 3. As α increases the glottal gap extends towards the anterior edge of the replica. It was shown [10] that analytical expressions for $A(\alpha)$, $l_A(\alpha)$ and $h_A(\alpha)$ can be derived from the other dimensions indicated in Fig. 3, i.e. largest E and smallest e VF thickness along the inferior–superior direction, length L along the posterior–anterior direction, constant distance $a = 4.5$ mm between the anterior VF’s edge and the rotation axis in the posterior–anterior direction.

Three contact regimes are then identified depending on the degree of contact ($1 - l_A(\alpha)/L$) between the medial surfaces of the left and right VF: full contact in regime I ($l_A = 0$), partial contact in regime II ($0 < l_A < L$) and no contact in regime III ($l_A \geq L$). Two critical angles $\alpha_{I,II}$ and $\alpha_{II,III}$ are then defined: $\alpha_{I,II}$ indicating the largest angle for which full contact occurs (shift from regime I to II) and $\alpha_{II,III}$ indicating the largest angle for which partial contact occurs (shift from regime II to III). Critical angles for each of the VF replicas are listed in Table 1. Angles are similar for the M5 and MRI replica and about half these values for the EPI replica.

2.3. Imposed α and relevance to UVFP

Angular asymmetry angles $\alpha \leq 20^\circ$ are experimentally imposed: $\alpha \in \{0^\circ, 4^\circ, 10^\circ, 16^\circ, 20^\circ\}$ for the M5 replica and $\alpha \in \{0^\circ, 4^\circ, 10^\circ, 20^\circ\}$ for the MRI and EPI replicas. Following Table 1, contact regime III (no contact) occurs only for the EPI VF replica at $\alpha = 20^\circ$ as $\alpha_{II,III} > 20^\circ$ holds for both the MRI and EPI VF replicas. Contact regime I

(full contact) and regime II (partial contact) are explored for all three replicas as indicated in Table 1.

In terms of UVFP, the vertical level difference between the superior transverse surfaces of the left and right VF is one of the sought clinical markers, reported to range up to 3 mm for UVFP subjects [5]. For each replica, two vertical level difference measures are considered, i.e. maximum vertical level difference $h_A(\alpha)$ and overall vertical level difference $\bar{h}(\alpha) = A(\alpha)/l_A(\alpha)$ so that $\bar{h}(\alpha) = h_A(\alpha)/2$. From Fig. 4 it can be seen that $h_A(\alpha) < 6$ mm and $\bar{h}(\alpha) < 3$ mm, respectively twice and once the range (up to 3 mm) reported for human subjects. It follows that measurements for $\alpha = 20^\circ$ fall either just outside (h_A) or just within (\bar{h}) the range observed on human subjects. This illustrates the need, reported [3], for a clear definition and precise measurement technology of vertical level difference in order to facilitate the comparison of findings on human subjects.

Regardless of their discrepancy, experimentally assessed maximum $h_A(\alpha)$ and overall $\bar{h}(\alpha)$ are both of the order of magnitude observed in human subjects. Consequently, assessed asymmetry angles α allow to investigate for each replica the influence of air leakage due solely to angular asymmetry in a range pertinent to UVFP. Inter-replica variations are then attributed to their different multi-layered structure and shape as outlined in Section 2.1.

3. Experimental setup

Each VF replica with imposed angular asymmetry α is mounted in the experimental setup depicted in Fig. 5 so that a fluid–structure interaction leads to auto-oscillation during which aerodynamic (Fig. 5(a)) and vibration (Fig. 5(b)) data are acquired. A rigid uniform tracheal tube (diameter 16 mm, length 460 mm, acoustic resonance frequency 185 Hz) is attached airtight to the inferior end of the glottal VF replica. Continuous steady airflow (density $\rho_G = 1.2$ kg m⁻³, dynamic viscosity $\mu_G = 1.8 \times 10^{-5}$ Pa s, temperature 24 ± 2 °C) is provided by an air compressor (Hitachi SC820) connected to a pressure reservoir (volume 0.04 m³). The pressure reservoir is filled with acoustic foam in order to avoid parasite acoustic resonances. The compressor is equipped with a pressure regulator (10202U, Fairchild, Winston-salem, NC). A pressure transducer (Kyowa PDS-70GA, accuracy ± 5 Pa) is positioned in a pressure tap in the tracheal wall, 185 mm upstream of the glottal VF replica, in order to measure upstream pressure P_u as a function of time t with sampling frequency 10 kHz. The setup described so far is similar to the one detailed in [10].

In addition to this, a high speed (HS) camera (Keyence VW-9000, frame rate 4 kHz, shutter time 4 μ s and resolution 240 px \times 320 px) is used to acquire instantaneous VF images, i.e. two dimensional greyscale intensity matrices $I(x, y, t)$ (or $I_{xy}(t)$ in short) with x and y indicating the pixel position as shown in Fig. 5(b). Image sequences are cropped from their original size 240 \times 320 to the region-of-interest containing the vibrating VF’s. It follows that in general $1 \leq x \leq N_x$ and $1 \leq y \leq N_y$ with N_x and N_y indicating the number of pixels in the x and y dimension, respectively. A single HS camera is rotated in the medio-frontal plane, i.e. around the posterior–anterior axis, to observe the VFs (distance 7 cm) from three distinct viewing angles θ_{HS} , taken

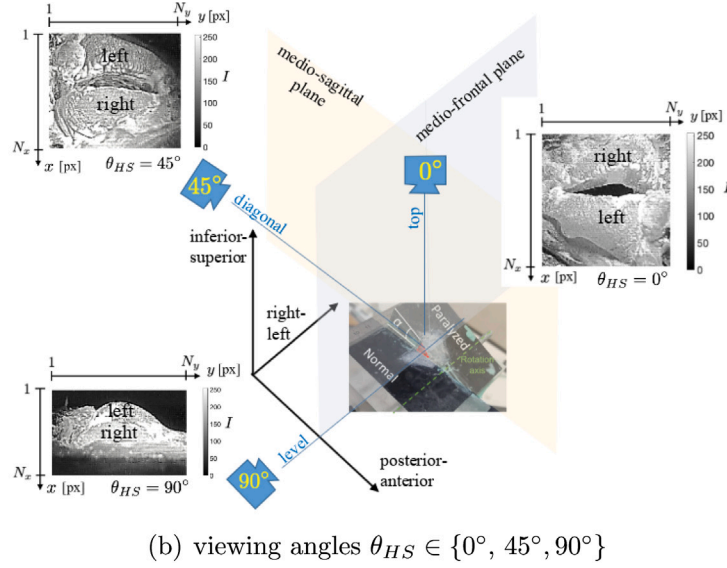
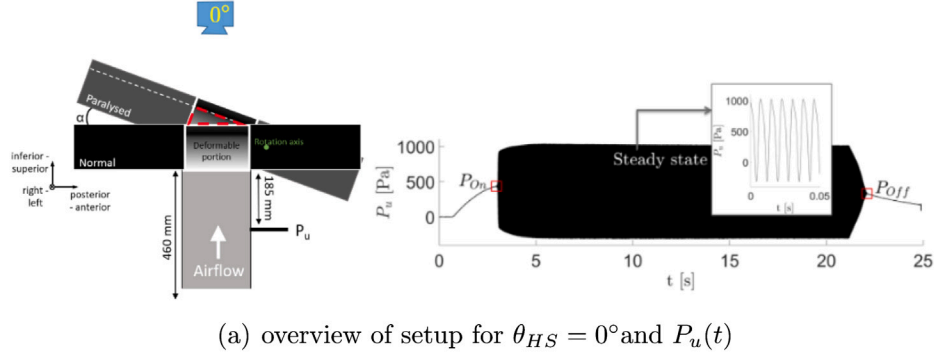


Fig. 5. (a) Overview of experimental setup and typical experiment: imposed asymmetry angle α , upstream pressure (P_u) sensor tap, HS camera (top view $\theta_{HS} = 0^\circ$) and $P_u(t)$ indicating auto-oscillation onset, steady state and offset. (b) HS camera viewing angles: $\theta_{HS} = 0^\circ$ (top view), $\theta_{HS} = 45^\circ$ (diagonal view) and $\theta_{HS} = 90^\circ$ (level view).

as the angle between the observation direction and the medio-sagittal plane, depicted in Fig. 5(b): top view $\theta_{HS} = 0^\circ$, horizontal level view $\theta_{HS} = 90^\circ$ and diagonal view $\theta_{HS} = 45^\circ$.

The fluid–structure interaction leading to auto-oscillation is illustrated plotting $P_u(t)$ in Fig. 5(a). As airflow is supplied, upstream pressure P_u gradually increases until auto-oscillation starts as P_u reaches the onset pressure threshold for auto-oscillation P_{On} . Steady state auto-oscillation is then observed for several seconds during which mean upstream pressure $\bar{P}_u \approx P_{On}$. Finally, auto-oscillation ceases as P_u is reduced below the offset pressure threshold for auto-oscillation P_{Off} . In the following, vibration motion dynamics for each VF replica condition is assessed on 1 s (corresponding to 4000 subsequent images) of the steady state auto-oscillation indicated in Fig. 5(a).

4. Image analysis

4.1. Global: image correlation

For each viewing angle θ_{HS} , the image correlation coefficient $\gamma(t)$ between image $I_{xy}(t)$ and reference image $J_{xy} = I_{xy}(t_{ref})$ is calculated. Consequently, an image-based correlation (IC) time series $\gamma(t)$ is obtained quantifying the instantaneous degree ($|\gamma| \leq 1$) of similarity to J_{xy} as:

$$\gamma(t) = \frac{\sum_x \sum_y (I_{xy}(t) - \bar{I})(J_{xy} - \bar{J})}{\sqrt{\left(\sum_x \sum_y (I_{xy}(t) - \bar{I})^2\right) \left(\sum_x \sum_y (J_{xy} - \bar{J})^2\right)}}, \quad (1)$$

where averaged image intensities are obtained as $\bar{I} = \sum_{xy} I_{xy}/N$ and $\bar{J} = \sum_{xy} J_{xy}/N$ with $N = N_x \times N_y$, the number of pixels in each image.

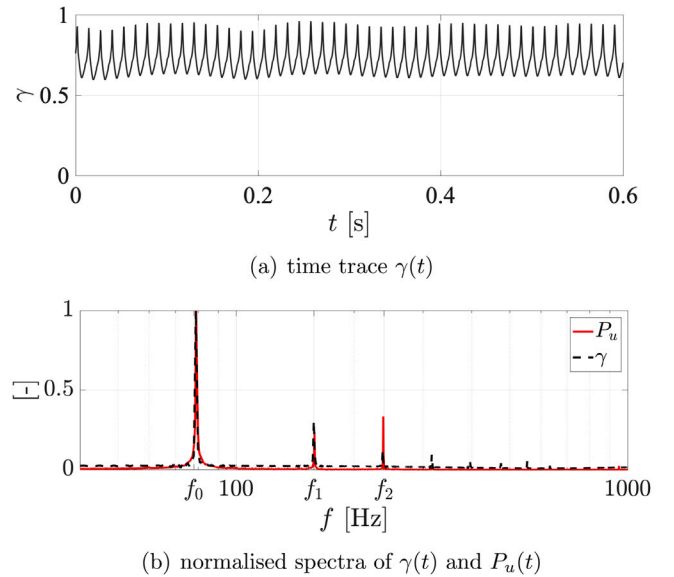


Fig. 6. Illustration for the EPI VF replica with $\alpha = 20^\circ$ and $\theta_{HS} = 90^\circ$ of: (a) image-based correlation $\gamma(t)$, (b) Normalised amplitude spectra from $\gamma(t)$ (dashed line) and from upstream pressure $P_u(t)$ (full line) and harmonic frequencies $f_{0,1,2}$. The frequency scale is logarithmic.

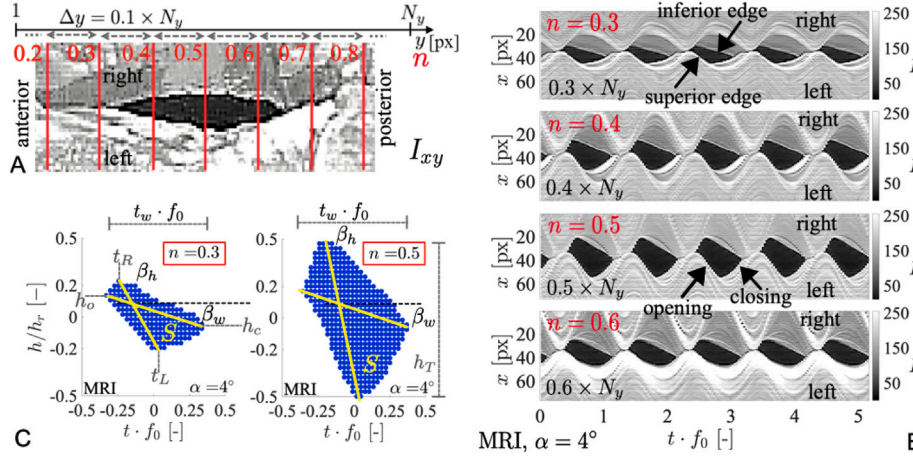


Fig. 7. Illustration of VK analysis for top view images $\theta_{HS} = 0^\circ$: line extraction of instantaneous image I_{xy} for $y = n \times N_y$ (frame A), 5 periods of each multi-VK image set (frame B) and feature extraction on mean aperture (frame C).

A time trace of $\gamma(t)$ is illustrated in Fig. 6(a) (EPI replica, $\alpha = 20^\circ$, $\theta_{HS} = 90^\circ$).

Normalised amplitude spectra of $\gamma(t)$ are considered from which oscillation frequency f_0 is derived. As an example, IC (from $\gamma(t)$) and aerodynamic (from $P_u(t)$) amplitude spectra are plotted in Fig. 6(b) (EPI replica, $\alpha = 20^\circ$, $\theta_{HS} = 90^\circ$). Harmonic frequencies f_1 and f_2 are indicated.

4.2. Local: left–right videokymographic vibration parameters

High-speed videokymography (VK) is a common technique to observe the VF's vibration dynamics from top view imaging ($\theta_{HS} = 0^\circ$) on human subjects [19] as well as on VF replicas [14]. Whereas the proposed IC analysis Eq. (1) results in a global image feature $\gamma(t)$, VK allows a local analysis as it scans in each image $I_{xy}(t)$ along the right–left x -direction pixels with constant posterior–anterior y -position. As tilting ($\alpha > 0^\circ$) of the left VF might provoke an asymmetry in the displacement of each single VF along the y -axis, line extraction in I_{xy} is done for several y -positions n so that $y = n \times N_y$, with $n \in \{0.2, \dots, 0.8\}$ as indicated by the vertical lines (red) in part A of Fig. 7. It follows that the line-scan at $y = 0.2 \times N_y$ is near the anterior edge and the rotation axis. The line-scan at $y = 0.8 \times N_y$ is near the posterior edge where the glottal gap emerges for $\alpha > \alpha_{(I,II)}$ as outlined in Section 2.2. Consecutive line scans in time result in a set of VK images associated with each y -position as illustrated in frame B of Fig. 7. Shown VK images allow to visualise the displacement of the left and right VF towards and away from each other during the closing and opening portion of the glottal cycle. It is noted that for $\alpha > \alpha_{I,II}$ apparent full closure in the VK images might mask air leakage due to the glottal gap in the medio-sagittal plane.

The analysis of each VK image relies on a common threshold-based segmentation of the VF edges and aperture for each cycle. The cycle's mean aperture is determined as the set of pixels surrounding the centroids recurring in at least 80% of individual cycles. As such each VK image condition (replica, angular asymmetry α and position n) is characterised by its mean aperture and its edges visualising the closing and opening of the left and right VF at that position. Resulting mean apertures for $n = 0.3$ and $n = 0.5$ are illustrated in frame C of Fig. 7. Time t is normalised by the oscillation period $1/f_0$ so that each normalised cycle has unit period. The displacement from the centroid h is normalised by the maximum aperture amplitude h_r for all n so that $-0.5 \leq h/h_r \leq 0.5$. These normalisations facilitate comparison between different conditions determined by the used replica, α and position n .

Glottal aperture parameters and left–right vibration asymmetry are quantified on the mean aperture and its extrema illustrated in frame C of Fig. 7.

4.2.1. Glottal aperture measures

The glottal aperture waveform is characterised by its area S . The open quotient is given by the duration of the open portion $t_w \cdot f_0 \leq 1$ as the normalised cycle period has unit length. It follows that the closed portion in the VK image has duration $1 - t_w \cdot f_0$. The skewness of the aperture is expressed considering angles β_w and β_h with respect to the direction of the normalised time axis. Angle β_w is determined by the diagonal direction associated with aperture extrema along the time axis. Hence $\beta_w \neq 0^\circ$ indicates asymmetry in the displacement of the left and right VF at the beginning (h_o) and end (h_c) of the open portion in each VK image. Angle β_h is determined by the diagonal direction associated with aperture extrema along the displacement axis h/h_r . Hence $\beta_h \neq 90^\circ$ indicates asymmetry with respect to the timing at which the displacement of inferior L and R fold edges becomes smaller than the displacement of their respective superior edges.

4.2.2. Left–right vibration asymmetry parameters

The angle β_h is related to the left–right phase asymmetry PA [21], which quantifies the normalised phase delay between the left and right VFs as,

$$PA = \frac{t_L - t_R}{t_w \cdot f_0}. \quad (2)$$

The angle β_w is related to the axis shift AS quantifying the normalised displacement of the VF during the VK's closed cycle portion [21] as,

$$AS = \frac{h_o - h_c}{h_T}, \quad (3)$$

with h_T the maximal aperture amplitude.

4.2.3. Left–right mucosal wave asymmetry

Extreme positions on the mean aperture associated with $\{t_L, t_R, h_o, h_c\}$ in Fig. 7 allow to segment the superior and inferior edges of the right and left VF during the opening (increasing $|h/h_r|$) and closing (decreasing $|h/h_r|$) portion of the cycle respectively. Noting normalised time $u_i = t_i \cdot f_0$ and normalised edge displacement $s = h/h_r$, the wave parameters are determined approximating each edge portion $s(u_i)$ by a Fourier series expansion with M harmonics,

$$s(u_i) = B_0 + \sum_{k=1}^M [A_k \sin(2\pi k u_i) + B_k \cos(2\pi k u_i)], \quad (4)$$

where coefficients B_0 , A_k and B_k are estimated using a least square fit [22]. It follows that each mucosal edge wave is described as

$$s(u_i) = B_0 + \sum_{k=1}^M [C_k \cos(2\pi k u_i + \phi_k)], \quad (5)$$

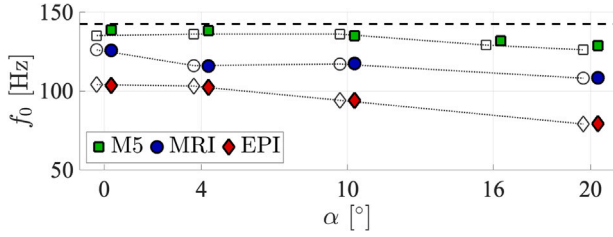


Fig. 8. Oscillation frequency $f_0(\alpha)$ for VF replicas M5 (\square), MRI (\circ) and EPI (\diamond) from aerodynamic $P_u(t)$ (empty symbols) and IC $\gamma(t)$ for $\theta_{HS} = 90^\circ$ (filled symbols) shifted around α and mechanical resonance frequency $f_1^M = 142$ Hz (horizontal dashed line).

with amplitude C_k and phase ϕ_k of the k th harmonic given as

$$C_k = \sqrt{A_k^2 + B_k^2}, \quad (6)$$

$$\phi_k = -\tan^{-1}\left(\frac{A_k}{B_k}\right). \quad (7)$$

Following [22] the first-order vibration mode ($M = 1$) was used to estimate the wave parameters for each edge. The phases of the inferior and superior edges of the right VF are denoted $\phi_{R,I}$ and $\phi_{R,S}$, respectively. Similarly, phases for the left VF are denoted $\phi_{L,I}$ and $\phi_{L,S}$.

The phase difference between the superior and inferior edge [23,24] for the right and left VF yields

$$\phi_R = \phi_{R,S} - \phi_{R,I}, \quad (8)$$

$$\phi_L = \phi_{L,S} - \phi_{L,I}. \quad (9)$$

The phase difference is related to the mucosal wave velocity c_R and c_L of the right and left VF as

$$\phi_{R(L)} = \frac{2\pi z_{R(L)}}{c_{R(L)}}, \quad (10)$$

with z_R and z_L the distance along the inferior–superior direction between the inferior and superior edges of the right (z_R) and left (z_L) VF. Note that thanks to the applied time normalisation the cycle has unit period so that Eq. (10) does not depend on frequency. Consequently, when $z_R \approx z_L$, using the assumption that the VF shape is not affected by α , the left–right wave velocity asymmetry is given by the left–right asymmetry in phase difference

$$\frac{\phi_R}{\phi_L} \approx \frac{c_L}{c_R}. \quad (11)$$

5. Results

5.1. Global analysis

Oscillation frequencies $f_0(\alpha)$ quantified from aerodynamic data $P_u(t)$ (empty symbols) and image-based correlation IC $\gamma(t)$ for $\theta_{HS} = 90^\circ$ (filled symbols) match for all assessed conditions as illustrated in Fig. 8. The maximum discrepancy between different viewing angles θ_{HS} yields less than 4 Hz, which corresponds to the experimental accuracy related to repeatability. Firstly, this confirms that f_0 can be obtained accurately from vibration imaging using the time series $\gamma(t)$ proposed in Eq. (1). Secondly, it shows that f_0 quantified from IC $\gamma(t)$ is independent from the camera viewing angle θ_{HS} . It is noted from Eq. (1) that the choice of reference image J_{xy} in the image sequence does not alter the time-dependence of $\gamma(t)$ and hence obtained f_0 .

From Fig. 8 is then seen that both aerodynamic $P_u(t)$ and IC $\gamma(t)$ time series reveal the decrease of oscillation frequency f_0 with increasing α confirming previous observations [10,14]. The magnitude of the decrease depends on α as well as on the used VF replica, i.e. $\leq 8\%$ for M5, $\leq 14\%$ for MRI and $\leq 25\%$ for EPI, expressing the influence of replica composition and geometry on the vibration.

Table 2
Angular asymmetry classification for $\alpha > 0^\circ$ for VF replicas.

| | mild | moderate | severe |
|-----|-----------|------------|----------------------|
| M5 | 4° | 10° | $16^\circ, 20^\circ$ |
| MRI | 4° | 10° | 20° |
| EPI | – | 4° | $10^\circ, 20^\circ$ |

It was shown on aerodynamic data $P_u(t)$ that increasing α is associated with an increased contribution of higher harmonic frequencies to the time trace [10]. Nevertheless, as peak values associated with harmonic frequencies for image-based correlation spectra are found to depend on the imposed camera view angle θ_{HS} the harmonic content of $\gamma(t)$ is not further quantified. Consequently, also harmonic peak values of aerodynamic and correlation spectra exhibit different harmonic amplitudes, e.g. for f_2 in Fig. 6(b).

5.2. Local analysis

Vibration observation from VK images is illustrated in Appendix. The influence of α is easily seen for the MRI replica in Fig. A.16. The impact of the used replica is apparent from Fig. A.17 for both $\alpha = 0^\circ$ and $\alpha = 20^\circ$. The impact of scan position n is shown in frame B of Fig. 7 for the MRI VF replica with $\alpha = 4^\circ$. In the following, measures defined in Section 4.2 are quantified and it is sought to which extent they capture qualitative observations with respect to the used replica (and hence its structure and geometry), scan position n and degree of angular asymmetry α . For convenience and considering the vertical level differences plotted in Fig. 4 and critical angles given in Table 1, the angular asymmetry is labelled *mild* for $0 < \alpha \leq \alpha_{I,II}$, *moderate* for $\alpha_{I,II} < \alpha \leq \alpha_{II,III}/2$ and *severe* for $\alpha_{II,III}/2 < \alpha$. The resulting angle classification for each replica is summarised in Table 2.

5.2.1. Glottal aperture measures

The normalised aperture area $0 < S/S_r \leq 1$ is plotted in Fig. 9 for each VF replica where the reference area S_r is set to its overall maximum. For all replicas, varying α between 0° and 20° alters S/S_r up to 80% depending on n . Indeed, varying n also induces S/S_r to differ up to 80% depending on the imposed α as in general S increases with n until a maximum is reached after which S reduces. The position of this maximum is more posterior (n increased) for the MRI replica compared to the EPI and M5 replica where the maximum is centred, showing the influence of the replicas geometry. Furthermore, S decreases with α for the M5 replica whereas it increases for MRI and EPI suggesting the influence of the replica composition. The increase with α is gradual for M5. For MRI and EPI the decrease is more pronounced for severe angular asymmetry.

The open quotient $t_w \cdot f_0 < 1$ is plotted in Fig. 10. Tendencies agree mostly with those reported for area S/S_r indicating that overall larger aperture areas correspond to longer open portions. However, the dependence on position n is less pronounced for $0.3 < n < 0.8$ as the open quotient is fairly constant for mild and moderate angular symmetry in the case of the EPI and MRI replicas and remains constant for angular asymmetry levels ($\alpha > 0$) in the case of the M5 replica. Consequently, open quotient $t_w \cdot f_0$ is more robust than S/S_r with respect to scan position n whereas it is less sensitive to the degree of angular asymmetry α and the replica's geometry. It is noted for MRI and EPI that the reduction of open quotient $t_w \cdot f_0$ with α is naturally associated with a reduction of the useful scan region along the posterior–anterior direction.

Aperture skewness angles β_h and β_w are plotted in Fig. 11 and Fig. 12, respectively. Without angular asymmetry ($\alpha = 0^\circ$), β_h and β_w approximate $\beta_h \approx 90^\circ$ and $\beta_w \approx 0^\circ$ at centre positions $0.3 < n < 0.7$ expressing symmetrical left–right VF displacement. Imposing angular asymmetry ($\alpha > 0^\circ$) induces skewness as β_h increases with 10° and β_w decreases with 10° for all replicas. Note that S/S_r and $t_w \cdot f_0$ exhibit

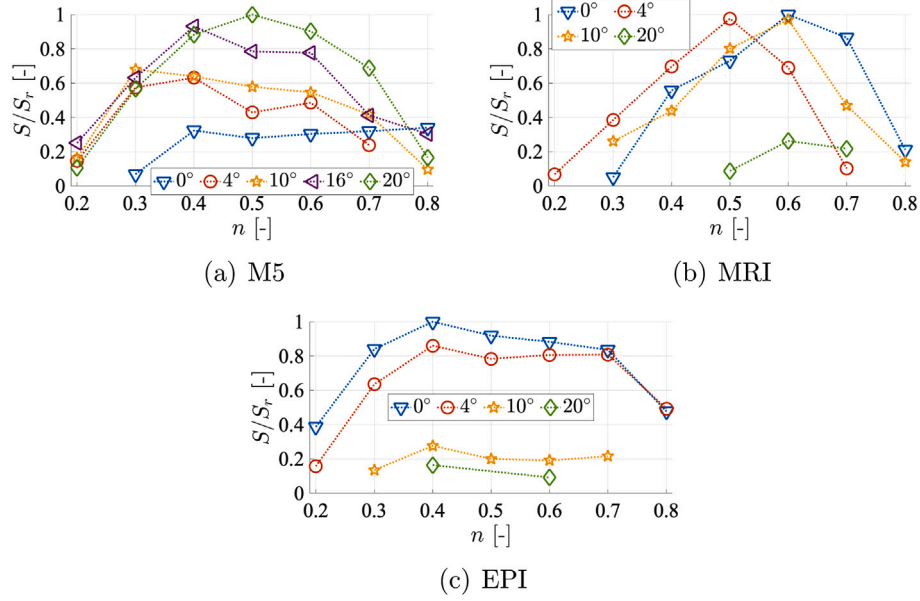


Fig. 9. Normalised aperture area $S(\alpha)/S_r$ at different positions n for replicas: (a) M5, (b) MRI and (c) EPI.

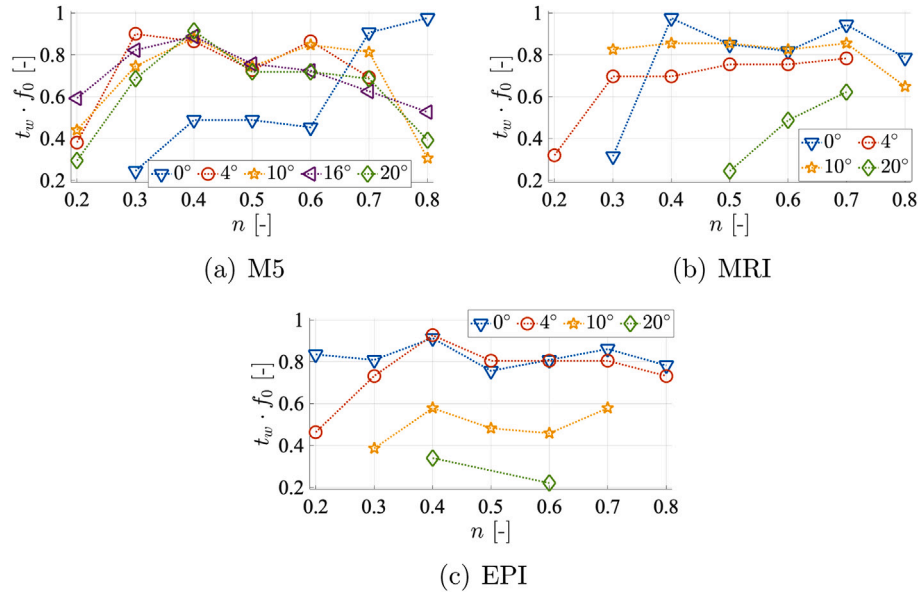


Fig. 10. Open quotient $t_w \cdot f_0$ as a function of α at different positions n for replicas: (a) M5, (b) MRI and (c) EPI.

opposite tendencies with α , *i.e.* increase for M5 and decrease for MRI and EPI. It follows that both β_h and β_w reveal angular asymmetry for $\alpha > 0$ indicating that left–right VF displacement asymmetry and subsequent skewness are inherent features of angular asymmetry. In general, skewness angles β_h and β_w do not reflect the severeness of angular asymmetry or scan position n as no general monotonic tendencies with α are distinguished. Nevertheless for β_w it is noted that such a tendency is observed for the M5 replica as β_w gradually decreases with both α and n , so that β_w is more prominent for positions near the posterior side which is opposite from the rotation axis. For the EPI replica (and for MRI at large n) the monotone decrease with α ceases when $\alpha = 20^\circ$. This is probably due to the small aperture area (Fig. 9) and associated open quotient (Fig. 10) at $\alpha = 20^\circ$, limiting the decrease for β_h and β_w .

5.2.2. Left–right vibration asymmetry

Following their definitions, left–right phase asymmetry PA (Eq. (2)) is related to β_h and axis shift AS (Eq. (3)) is related to β_w . Phase

asymmetry PA increases from its symmetry value $PA = 0$ with α . Observed tendencies for PA are similar to the ones described for β_h so that PA curves are not plotted. It follows that left–right phase asymmetry can be quantified by either PA or β_h . As $PA > 0$ (or equivalently $\beta_h > 90^\circ$) holds, this implies that the right VF reaches its maximum displacement followed by the tilted left VF. This agrees with the observation during auto-oscillation experiments that the movement of the right VF entrains the movement of the tilted left VF.

From Fig. 13 it can be seen that for $\alpha > 0$ axis shift AS increases from its symmetry value $AS = 0$ approximated for $\alpha = 0^\circ$ for all replicas. For the EPI and MRI replicas findings for β_w also apply to AS so that AS is related to the severeness of angular asymmetry up to $\alpha = 20^\circ$. For the M5 replica the dependence on n observed for β_w is also retrieved for AS . Nevertheless, AS does no longer relate to the severeness of angular asymmetry as values for moderate and severe asymmetry overlap. For $\alpha > 0^\circ$, axis shift $AS > 0$ (or $\beta_w < 0^\circ$) indicates left–right asymmetrical mucosal wave displacement during the closed

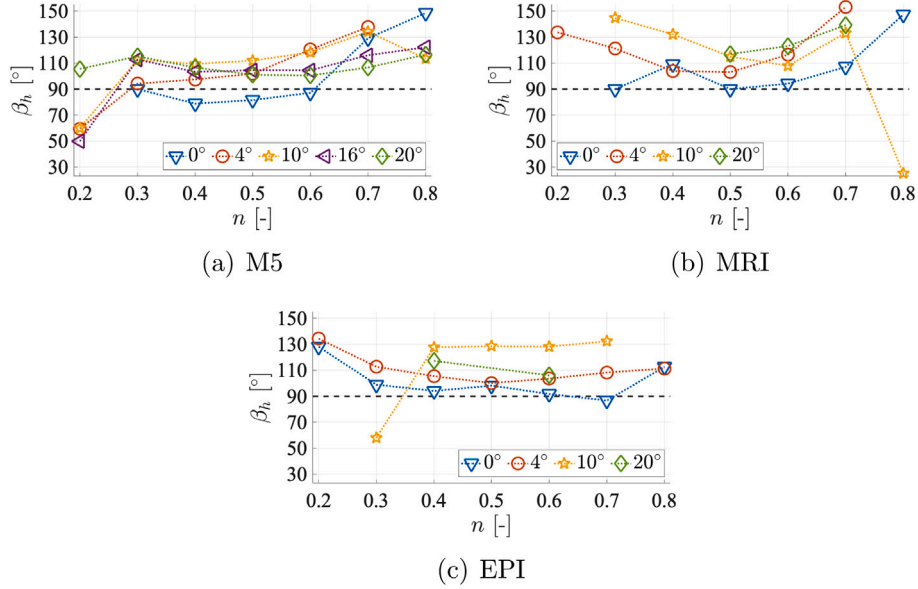


Fig. 11. Aperture angle $\beta_h(\alpha)$ at different positions n for replicas: (a) M5, (b) MRI and (c) EPI. The symmetry condition $\beta_h = 90^\circ$ is indicated (dashed line).

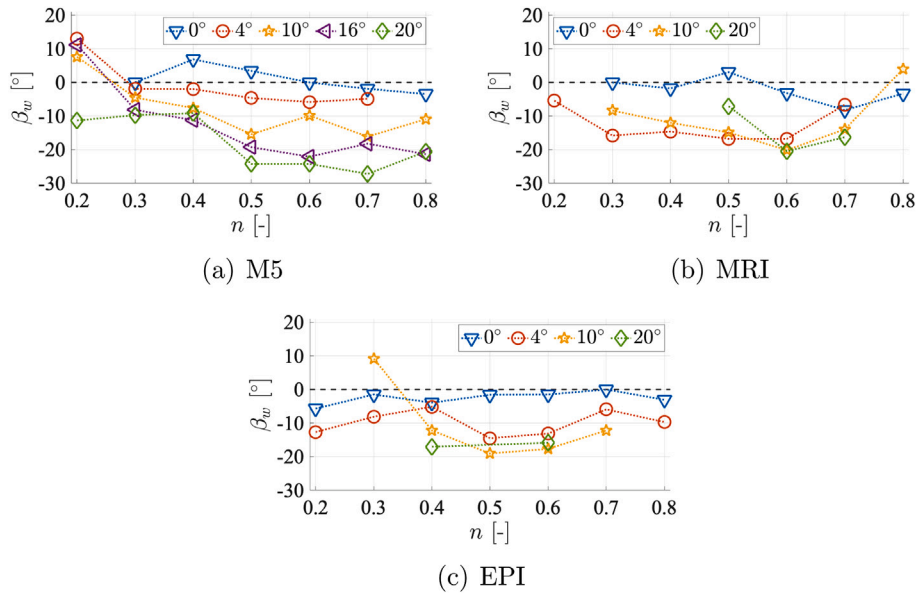


Fig. 12. Aperture angle $\beta_w(\alpha)$ at different positions n for replicas: (a) M5, (b) MRI and (c) EPI. The symmetry condition $\beta_w = 0^\circ$ is indicated (dashed line).

portion. The mucosal wave asymmetry is further assessed in the next section.

5.2.3. Left-right mucosal wave asymmetry

The mucosal wave phase difference between the superior and inferior edges of the left VF ϕ_L (Eq. (9)) normalised by $\phi_{L,r}$, taken as the mean phase difference for all n at symmetry ($\alpha = 0^\circ$), is illustrated in Fig. 14 for the M5 and EPI replicas. It is found that for the MRI and EPI replicas ϕ_L increases with α . The increase is most prominent for moderate and severe asymmetry angles. For the M5 replica ϕ_L decreases when $\alpha > 0^\circ$ and values for $\alpha > 0^\circ$ overlap. The phase difference of the right VF ϕ_R is not shown as tendencies are similar as those observed for ϕ_L .

The phase difference ratio between the right and left VF ϕ_R/ϕ_L is plotted in Fig. 15. The ratio also informs on the left-right wave velocity asymmetry c_L/c_R using Eq. (10). In general, the symmetry condition $\phi_R/\phi_L = 1$ holds for $\alpha = 0^\circ$ as phase differences and wave

velocities are equal for the left and right VF as expected. For the MRI and EPI replicas the ratio decreases (up to 60%) for mild and moderate asymmetry angles. For severe asymmetry, the ratio increases again towards (MRI) or slightly (<10%) above (EPI) unity. For the M5 replica a decrease of ϕ_R/ϕ_L with α occurs for moderate angular asymmetries. For severe asymmetries, ϕ_R/ϕ_L decreases (up to 80%) for locations near the anterior axis, whereas they remain around unity for locations near the posterior axis. For mild asymmetries, the ratio ϕ_R/ϕ_L also remains near unity. This implies that in general (when $\phi_R/\phi_L < 1$ holds) the phase difference is greater for the tilted left VF than for the right VF as $\phi_L \geq \phi_R$ and therefore $c_R \geq c_L$ so that the wave velocity associated with the tilted left VF is lower than the one in the right VF. This is consistent with the observations for PA (and therefore β_h), suggesting that the movement of the right VF entrains the left VF; and consistent also with the observation of axis shift AS (and therefore β_w) during the closed portion of the cycle. For the M5 replica $\phi_R/\phi_L \approx 1$ at scan position $n = 0.5$ for all α and for the MRI replica the overall

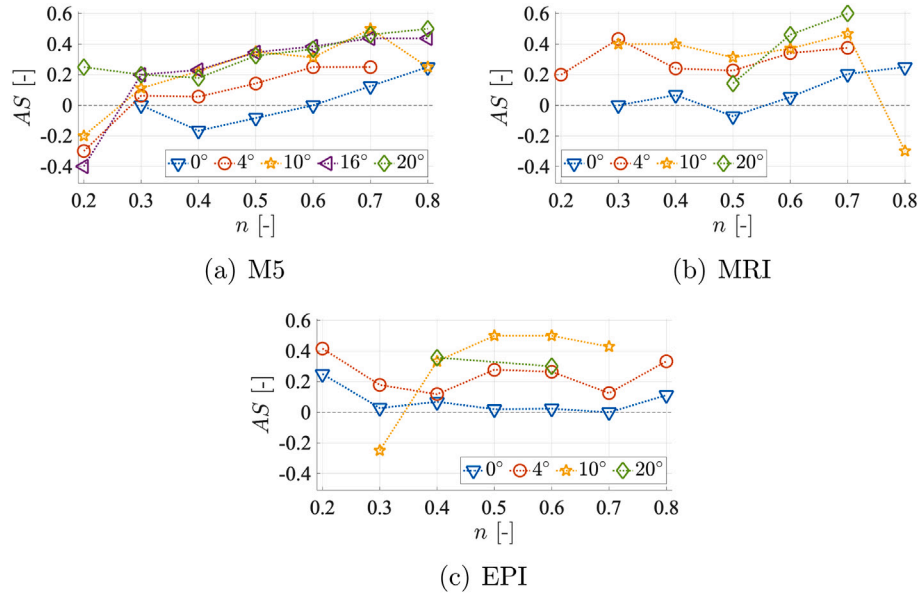


Fig. 13. Axis shift AS as a function of α at different positions n for replicas: (a) M5, (b) MRI and (c) EPI. The symmetry condition $AS = 0$ is indicated (dashed line).

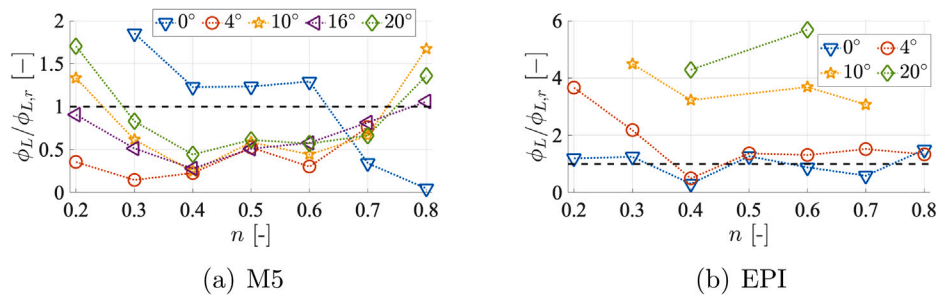


Fig. 14. Normalised phase difference of the left VF ($\phi_L/\phi_{L,r}$) as a function of α and position n for replicas: (a) M5, (b) EPI. Ratio 1 is shown (dashed line).

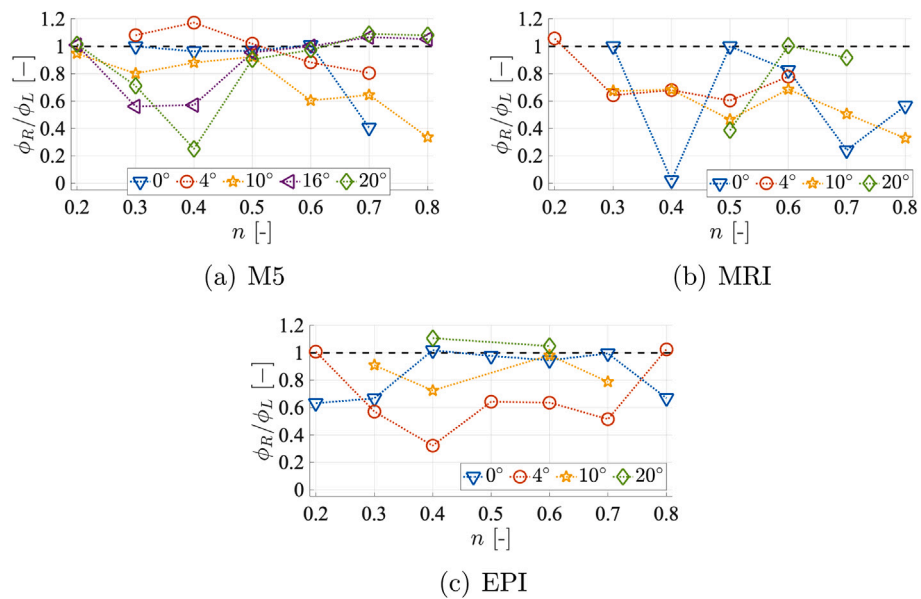


Fig. 15. Left-right phase difference asymmetry ϕ_R/ϕ_L as a function of α and position n for replicas: (a) M5, (b) MRI and (c) EPI. Phase symmetry $\phi_R/\phi_L = 1$ is indicated (dashed line).

discrepancy from unity is smallest at $n = 0.6$. It is suggested that these positions, where the discrepancy in left–right phase difference asymmetry with respect to α is smallest, indicate the entrainment position associated with synchronisation and hence the minimum in left–right phase difference asymmetry. The entrainment position is thus centred ($n = 0.5$) for the M5 replica and is shifted in the posterior direction ($n = 0.6$) due to the geometry of the MRI replica. Note that these positions correspond to maxima in the aperture area S/S_r . As the area variation of the EPI replica exhibits a broad maximum also the left–right phase difference asymmetry exhibits broader minimum discrepancy for $0.4 < n < 0.7$ corresponding to the entrainment region. As ϕ_R/ϕ_L depends on scan position n , a wave-like displacement along the posterior–anterior direction can be observed.

6. Discussion

The influence of air leakage due to vertical level difference created by imposing angular asymmetry between the right and tilted left VF is experimentally assessed using high-speed imaging during steady-state auto-oscillation of three deformable silicone VF replicas of different multi-layer composition and geometry. Observations for the MRI and EPI replicas are most relevant to human VF's and hence clinical applications as their structure and shape are based on human VF's. Nevertheless, it is of interest to study and compare vibration observations for all three replicas (M5, MRI and EPI) to understand their differences and similarities.

Image sequences are analysed globally introducing an image-based correlation IC time series $\gamma(t)$ from which the oscillation frequency is obtained regardless of degree of asymmetry, used replica or camera view angle. The shown robustness suggests that $\gamma(t)$ can be applied to clinical imaging data in order to obtain the oscillation frequency. Obtained oscillation frequencies match with frequencies retrieved on the upstream driving pressure $P_u(t)$, confirming the use of aerodynamic data to study vibration during auto-oscillation, and confirming the overall decrease of $f_0(\alpha)$ reported in [10]. This is in particular of interest for physical studies during which $P_u(t)$ acquisition is often more beneficial than imaging as it is readily accessible and does not demand a direct view of the vibrating VF's, reduces the amount of data storage and hence the amount of data processing and does not involve the use of (expensive) high-speed camera equipment.

Videokymographic images are analysed for different local scan positions n considering mean cycle apertures. To facilitate comparison between different conditions, the oscillation frequencies retrieved from the global image analysis are used for time normalisation and the displacement is normalised by the maximum cycle displacement. The influence of angular asymmetry, scan position and replica on left–right VF displacement asymmetry and mucosal wave asymmetry is quantified from common features relevant for clinical studies and for further studies on replicas.

Aperture's area S/S_r , open quotient $t_w \cdot f_0$ and superior versus inferior VF edge phase differences ϕ_L and ϕ_R reflects the replica's multi-layer composition as these quantities have opposite tendencies for the M5 replica on one hand and the softer MRI and EPI replicas on the other hand. This observation might be related to the minimum in the oscillation onset pressure threshold at $\alpha \approx 10^\circ$ found only for the M5 replica [10] indicating that for the M5 replica imposing angular asymmetry at first eases auto-oscillation. For the MRI and EPI replicas, these features can inform on angular asymmetry for severe asymmetries only as for values characterising no, mild and moderate asymmetry overlap. For clinical applications (and when a severe angular asymmetry is suspected) in particular the decrease of $t_w \cdot f_0$ is a potential indicator of severe angular asymmetry as the open quotient does not depend on scan position n and is known to vary between 0 and 1. Area S/S_r on the other hand is sensitive to scan position n and requires prior knowledge of a reference value S_r associated with symmetry ($\alpha = 0^\circ$) which might not be available for an individual human subject. It is

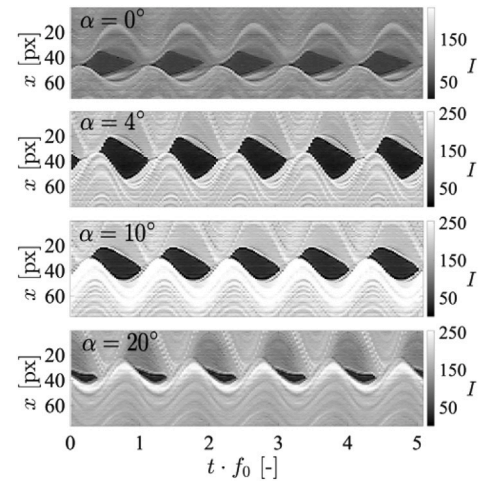


Fig. A.16. VK images for the MRI replica along line $y = 0.5 \times \Delta y$ (centre $0.5 \times N_y$, or $n = 0.5$) for increasing $\alpha = \{0^\circ, 4^\circ, 10^\circ, 20^\circ\}$ (top to bottom).

noted that for the M5 replica $t_w \cdot f_0$ only indicates the presence of angular asymmetry $\alpha > 0^\circ$ and not its severity. Since mild angular asymmetry might also occur for healthy VF's it seems not a pertinent parameter for the clinical study of angular asymmetry. In this case a combination with S/S_r is of interest in case reference value S_r is known since S/S_r relates to the degree of angular asymmetry for the M5 replica.

Aperture skewness angle $\beta_h > 90^\circ$ and phase difference $PA > 0$ both measure the increase of the phase asymmetry between the right and left VF with α . Aperture skewness angle $\beta_w < 0^\circ$ and axis shift $AS > 0$ indicate the increase of left–right asymmetrical mucosal wave displacement with α resulting in a decrease of the left–right phase difference asymmetry $\phi_R/\phi_L < 1$ with α . The tendencies observed for these asymmetry measures hold for all VF replicas so that they are identified as essential angular asymmetry parameters in contrast to previous discussed parameters (area S/S_r , open quotient $t_w \cdot f_0$ and phase differences ϕ_L and ϕ_R) for which tendencies also depend on the replica. Increasing phase difference (PA or β_h) indicates that angular asymmetry induces increased entrainment of the tilted left VF by the right VF. Decreasing β_w (or similarly increasing AS) indicates that angular asymmetry causes an increasingly asymmetric mucosal wave displacement during the closed portion between the left and right VF. Decreasing left–right phase difference asymmetry ϕ_R/ϕ_L indicates that angular asymmetry causes the mucosal wave velocity associated with the tilted left VF to lower with respect to the wave velocity of the right VF. It is noted that $\phi_R/\phi_L \approx 1$ or at least the decrease of $\phi_R/\phi_L \approx 1$ with α is limited for scan positions n within the entrainment region situated for all replicas near the centre position $n = 0.5$. It is noted that for the MRI replica with a more realistic shape entrainment is shifted in the posterior direction.

For the MRI and EPI replica phase these asymmetry measures vary gradually with angular asymmetry up to $\alpha < 20^\circ$ for which their value becomes limited due to the prominent area reduction. Therefore, for clinical applications it seems of interest to consider this parameter set (PA or equivalently β_h , AS , β_w and ϕ_R/ϕ_L) in combination with the open quotient $t_w \cdot f_0$ in order to assess the degree of angular asymmetry.

For future research on mechanical replicas it is of interest to verify that these findings hold for $\alpha < 0^\circ$ instead of $\alpha > 0^\circ$ as considered in the current study. Furthermore, described entrainment and vibration asymmetries are of interest for physical modelling of UVFP. Future clinical studies of UVFP are needed in order to consider a multi-variable analysis of the proposed asymmetry parameter set in order to assess that it allows firstly to detect the presence of UVFP and secondly allows to quantify the degree of UVFP. Note that in case this is the case purely

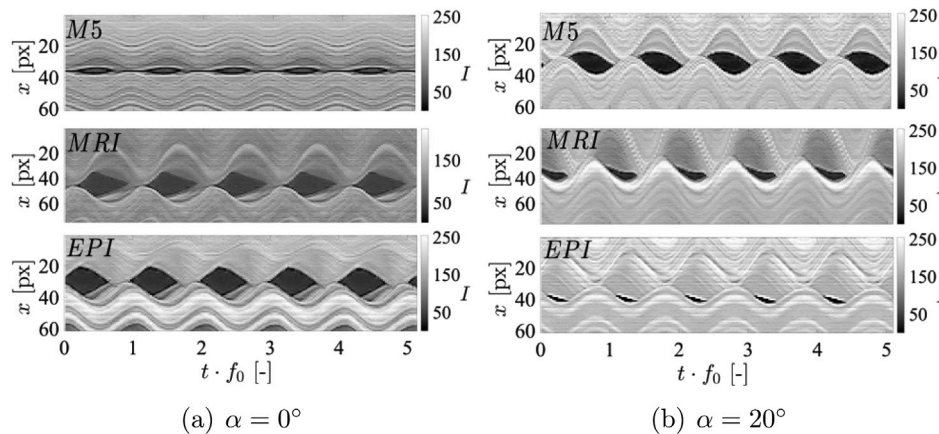


Fig. A.17. VK images for the M5, MRI and EPI replica (top to bottom) along line $y = 0.5 \times \Delta y$ (centre $0.5 \times N_y$ or $n = 0.5$) for: (a) $\alpha = 0^\circ$ and (b) $\alpha = 20^\circ$.

geometrical glottal changes, *i.e.* tilting, can at least partly explain UVFP symptoms.

7. Conclusion

An image-based analysis of steady-state auto-oscillation is presented using VF replicas with imposed angular asymmetry creating vertical level difference as observed in the case of UVFP. A global image-based correlation time series is shown to enable straightforward and robust estimation of the oscillation frequency for all assessed conditions, *i.e.* used replica, angular asymmetry and camera view angle. Videokymographic line scans are then analysed for several scan positions. Angular asymmetry is shown to induce left–right vibration phase asymmetry characterising the entrainment of the left VF by the right VF and a reduced mucosal wave velocity of the left VF compared to its velocity associated with the right VF. The extent of this vibration asymmetry induced by angular asymmetry is proposed to be quantified by a set of asymmetry parameters in combination with the open quotient. The use of different replicas has proven to be extremely valuable to identify these general mechanisms and associated parameter sets as general glottal aperture measures such as area and open quotient largely depend on the multi-layer composition of the used replica.

CRediT authorship contribution statement

Anne Bouvet: Concept of the study, Writing, Conducted the experiments. **Isao Tokuda:** Concept of the study, Writing. **Xavier Pelorson:** Concept of the study, Writing. **Annemie Van Hirtum:** Concept of the study, Writing, Analyses the data.

Declaration of competing interest

The authors declare that they have no known competing financial interests or personal relationships that could have appeared to influence the work reported in this paper.

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Appendix. Illustration of VK images

Examples of VK images illustrating the influence of imposed asymmetry angle α and the assessed replica, *i.e.* its shape and composition, on the vibration are shown in Fig. A.16 and Fig. A.17, respectively.

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