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An analytic model for the flow induced in syringomyelia cavities

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A simple two-dimensional fluid-structure-interaction problem, involving viscous oscillatory flow in a channel separated by an elastic membrane from a fluid-filled slender cavity, is analyzed to shed light on the flow dynamics pertaining to syringomyelia, a neurological disorder characterized by the appearance of a large tubular cavity (syrinx) within the spinal cord. The focus is on configurations in which the velocity induced in the cavity, representing the syrinx, is comparable to that found in the channel, representing the subarachnoid space surrounding the spinal cord, both flows being coupled through a linear elastic equation describing the membrane deformation. An asymptotic analysis for small stroke lengths leads to closed-form expressions for the leading-order oscillatory flow, and also for the stationary flow associated with the first-order corrections, the latter involving a steady distribution of transmembrane pressure. The magnitude of the induced flow is found to depend strongly on the frequency, with the result that for channel flow rates of non-sinusoidal waveform, as those found in the spinal canal, higher harmonics can dominate the sloshing motion in the cavity, in agreement with previous *in vivo* observations. Under some conditions, the cycle-averaged transmembrane pressure, also showing a marked dependence on the frequency, changes sign on increasing the cavity transverse dimension (i.e. orthogonal to the cord axis), underscoring the importance of cavity size in connection with the underlying hydrodynamics. The analytic results presented here can be instrumental in guiding future numerical investigations, needed to clarify the pathogenesis of syringomyelia cavities.

Key words:

1. Introduction

Syringomyelia is a condition characterized by the appearance of slender fluid-filled cavities, known as syrinxes, within the spinal cord (Rizk 2023). An illustration showing the typical location of the syrinx is given in figure 1(a). The condition frequently appears in patients with Chiari I malformation (Milhorat *et al.* 1999; George & Higginbotham 2011), a structural abnormality in which the lower part of the cerebellum herniates into the spinal canal, obstructing the normal flow of cerebrospinal fluid (CSF), the colorless

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39 Newtonian fluid that bathes the central nervous system. Alternative factors, such as
40 arachnoiditis, spinal cord tumors, or physical trauma, can also result in the formation of
41 a syrinx (Klekamp *et al.* 1997; Milhorat 2000).

42 The location of the syrinx within the spinal cord depends on the initiating cause.
43 For example, in syringomyelia linked to Chiari I malformation, syrinx cavities typically
44 form in the cervical region of the spine as an expansion of the central canal (*canalicular*
45 *syringomyelia*), a CSF-filled space that extends along the spinal cord (see figures 1(*b*)
46 and 1(*d*)). In contrast, for syringomyelia associated with spinal-cord trauma (*post-*
47 *traumatic syringomyelia*), *extracanalicular* syrinxes generally develop adjacent to the site
48 of the injury (Bertram 2009). The two types of syrinxes are represented in figures 2(*a*)
49 (canalicular syringomyelia) and 2(*b*) (extracanalicular syringomyelia), with the former
50 plot depicting a Chiari I malformation (see, e.g. Brodbelt & Stoodley 2003; Ahuja *et al.*
51 2017; Vaquero *et al.* 2017, for related clinical images).

52 Despite extensive research, the pathophysiology of the disease remains unclear (Stoodley
53 2014). Numerous theories have been advanced over the years (Elliott *et al.* 2013).
54 Since the conditions and injuries that precede syringomyelia involve abnormalities in the
55 motion of CSF, it is now generally agreed that CSF flow and its associated pressure
56 variations play an important role in the formation and enlargement of the cavity, as first
57 hypothesized by Gardner & Angel (1959).

58 Magnetic resonance imaging (MRI) techniques have been instrumental in gaining
59 understanding of the CSF flow dynamics. It is now well established that CSF displays
60 an oscillatory motion in the subarachnoid space (SAS) surrounding the spinal cord, as
61 indicated in figure 1(*d*). The oscillatory velocities, with peak values on the order of a few
62 centimeters per second, are driven by the respiratory and cardiac cycles (Linninger
63 *et al.* 2016; Kelley & Thomas 2023), with the former being dominant in the lumbar region
64 (Gutiérrez-Montes *et al.* 2022) and the latter being dominant in the cervical region (Yildiz
65 *et al.* 2017, 2022), where most syrinxes are formed.

66 Oscillatory motion synchronized with the cardiac cycle has also been observed inside
67 large syrinxes, with associated velocities comparable to those found in the SAS (Brugières
68 *et al.* 2000; Lichtor *et al.* 2005). For instance, Vinje *et al.* (2018) measured peak velocities
69 of 3.6 and 2.0 cm/s in the SAS and syrinx of a patient with Chiari I malformation, with
70 the values decreasing to 2.7 and 1.5 cm/s after the cavity shrank following surgery. As
71 indicated in the schematic of figure 1(*c*), the motion in the syrinx displays a sloshing
72 character, with the internal fluid motion inducing cyclic variations of the cavity shape
73 that can be visualized using high-resolution dynamic MRI (Honey *et al.* 2017). This
74 fluid slosh and its associated pressure fluctuations exert on the surrounding spinal-cord
75 tissue a cyclic traction that may contribute to the enlargement of the cavity (Honey
76 *et al.* 2017). As revealed by PC MRI measurements (Vinje *et al.* 2018), the motion in the
77 syrinx displays multiple oscillations per cardiac cycle, an intriguing feature of the flow
78 resulting from the fluid-structure dynamical interactions taking place.

79 Central to the pathophysiology of syringomyelia is the physical mechanism that pro-
80 duces the accumulation of fluid within the syrinx (the so-called “filling mechanism”
81 Stoodley 2014), a key aspect of the problem that remains unclear despite significant
82 research efforts (Williams 1980; Klekamp 2002; Heiss *et al.* 2019; Bhadelia *et al.* 2023).
83 Early investigators (Gardner & Angel 1959; Williams 1969) postulated that CSF flows
84 into the syrinx from the fourth ventricle of the brain through the central canal as a result
85 of a dissociation in craniospinal pressure. These initial ideas could not explain, however,
86 the development of the syrinx in patients with an obstructed central canal, that being
87 the case in most adults (Ball & Dayan 1972; Williams 1990; Garcia-Ovejero *et al.* 2015).

88 Alternative theories on the onset of syringomyelia point at a deregulation of the

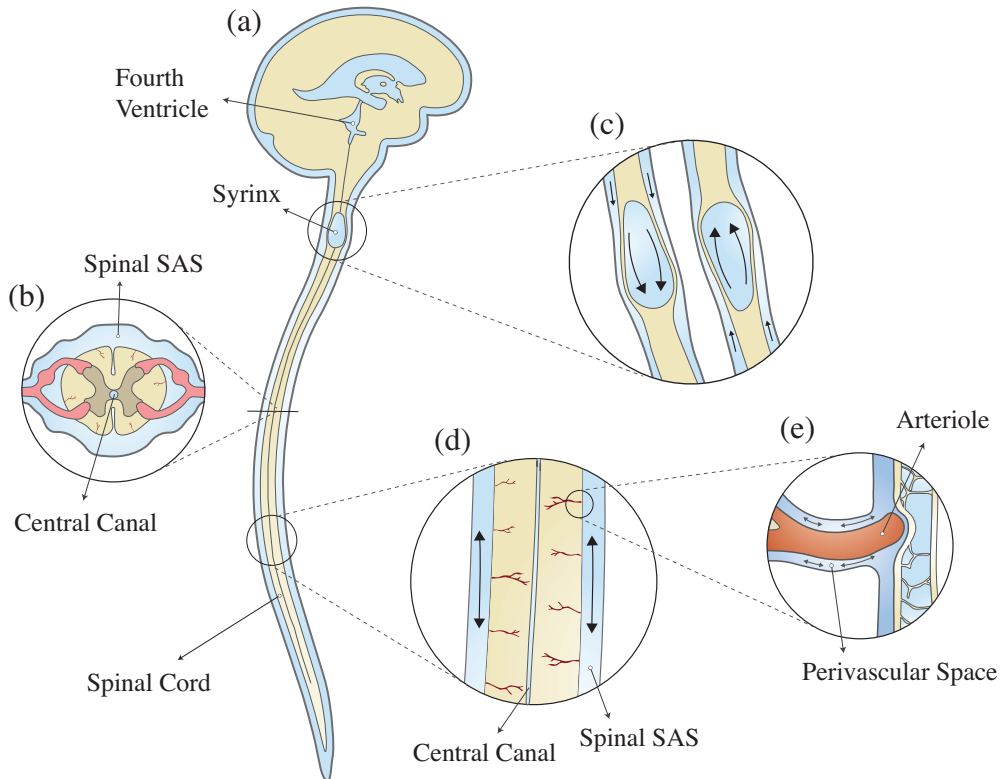


FIGURE 1. Schematic representation of the problem, including (a) a general view of the central nervous system for a subject having a syringomyelia syrinx at the cervical level, (b) view of the cross section of the spinal canal at a syrinx-free location, (c) a close view of the cavity with indication of the induced sloshing motion, (d) illustration of the longitudinal flow along the spinal SAS, and (e) close view of a spinal-cord periarterial space.

89 transmedullary flow established between the SAS and the central canal (Oldfield *et al.*
90 1994; Heiss *et al.* 1999, 2012; Lloyd *et al.* 2017; Heiss *et al.* 2019). *In vivo* experiments
91 using injection of fluorescent tracers in sheep, rats and mice have shown that radial inflow
92 and outflow occur predominantly along perivascular spaces surrounding blood vessels
93 (Stoodley *et al.* 1996, 1997; Wei *et al.* 2017; Liu *et al.* 2018, 2022). For instance, as shown
94 by Wei *et al.* (2017), when the tracer is released in the surrounding SAS, inflow occurs
95 mainly along the perivascular space surrounding penetrating arterioles (see figure 1(e)).
96 This phenomenon has been addressed by Bilston *et al.* (2010), who investigated effects of
97 changes in the timing of SAS pressure on perivascular flow into the spinal cord, and by
98 Elliott (2012), who developed one-dimensional models of transmedullary flow accounting
99 for the presence of perivascular spaces. Transmedullary tracer dispersion is assisted by
100 interstitial flow through the parenchyma (Wei *et al.* 2017), at different rates in grey and
101 white matter (Liu *et al.* 2018). The role of the spinal-cord-tissue poroelasticity in the
102 interstitial flow across the spinal cord has been investigated both numerically (Støverud
103 *et al.* 2016) and analytically (Cardillo & Camporeale 2021). An imbalance between the
104 inflow and outflow of CSF, associated with alterations of the transmedullary pressure
105 difference, may lead to accumulation of fluid within the cavity. In this regard, Ball &

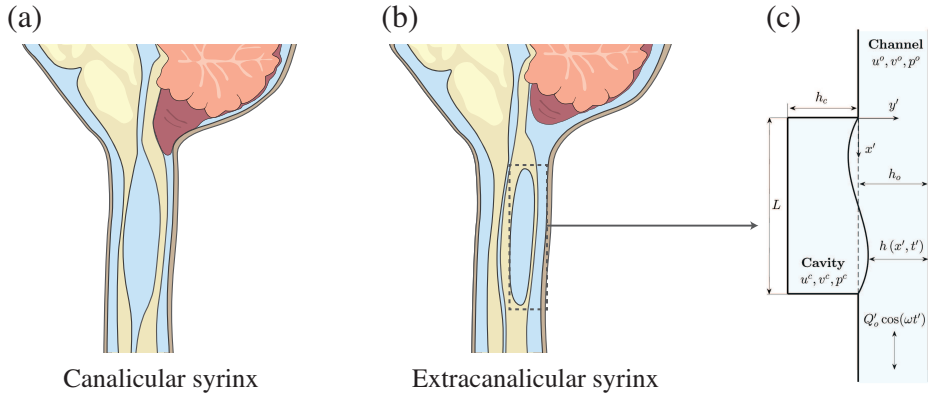


FIGURE 2. Schematic representations of *canalicular* (a) and *extracanalicular* (b) syringomyelia, and of the canonical model investigated here (c). The schematic (a) of the canalicular syrinx depicts a Chiari I malformation, while no specific cause is indicated for the extracanalicular case shown in (b).

106 Dayan (1972) suggest that sudden increases in thoracoabdominal pressure could force
 107 CSF into the cord, while Oldfield *et al.* (1994) argue that accentuated pressure waves
 108 transmitted by the downward displacement of the cerebellar tonsils during systole play a
 109 main role in syrinx formation. A key observation regarding syringomyelia is that the
 110 accumulation of fluid is very slow relative to the hydrodynamic time scales (Elliott
 111 *et al.* 2013), with the consequence that even quantitatively small changes of the existing
 112 pressure field, associated for instance with alterations of the normal CSF flow, may have
 113 a significant effect when acting over the long time scales characterizing cavity growth.

114 Numerical simulations and *in vitro* experiments have been extensively used to in-
 115 vestigate different aspects of syringomyelia hydrodynamics (Elliott *et al.* 2013). One-
 116 dimensional inviscid propagation of large-amplitude pressure waves along elastic channels
 117 was studied by Carpenter and coworkers (Berkouk *et al.* 2003; Carpenter *et al.* 2003) to
 118 ascertain whether the interactions of a large pressure impulse (e.g. generated by a cough
 119 or sneeze) with partial obstructions of the spinal canal could lead to damage of the cord
 120 tissue, a hypothesis not supported by subsequent studies (Bertram *et al.* 2005; Bertram
 121 2009; Elliott *et al.* 2009). The sloshing motion induced in the syrinx by a periodic pressure
 122 gradient has been investigated numerically (Bertram 2010; Drøsdal *et al.* 2013; Vinje
 123 *et al.* 2018) and experimentally (Martin *et al.* 2010). The studies of Bertram (2010) and
 124 Martin *et al.* (2010) considered a spinal cord with a large fluid-filled cavity adjacent to
 125 a SAS stenosis. As noted by Bertram (2010), the cycle-averaged pressure distribution
 126 resulting from fluid-structure interaction (FSI) involves a transmural pressure difference
 127 that could potentially drive CSF across the spinal SAS into the syrinx, a finding further
 128 corroborated in subsequent computations accounting for the permeability of the spinal
 129 cord (Heil & Bertram 2016; Bertram & Heil 2017).

130 Like the analyses mentioned in the preceding paragraph, the present paper addresses
 131 syringomyelia hydrodynamics, including the sloshing motion induced in the cavity by
 132 the oscillating SAS flow and the resulting transmural pressure. Unlike the previous
 133 investigations, however, our study is fundamentally analytic in nature, the aim being to
 134 clarify the essential FSI dynamics of syringomyelia cavities with use of a simple canonical
 135 model problem that affords description of elastic interactions between a confined fluid
 136 space and an open canal with oscillatory flow. In some sense, our approach is similar to

137 that followed in investigating oscillations in collapsible tubes (Grotberg & Jensen 2001;
138 Heil & Hazel 2011), for which the simple Starling resistor (Knowlton & Starling 1912)
139 is used as idealized canonical representation of the flow. Both planar and axisymmetric
140 configurations have been employed (Heil & Hazel 2011). The former, often used in Navier-
141 Stokes simulations of the flow (Heil & Hazel 2011), consists of a two-dimensional (2D)
142 channel in which a finite section of one of the rigid walls is replaced by a deformable wall,
143 represented by a prestressed elastic membrane that separates the channel fluid from a
144 pressure chamber. Wall deformations are induced by the viscous pressure variations in
145 the channel flow, with the wall stiffness dominated by the axial tension of the membrane,
146 leading to complicated FSI dynamical behaviors (Grotberg & Jensen 2001; Heil & Hazel
147 2011).

148 As shown in figure 2(c), the present analysis employs a variant of the 2D Starling
149 resistor to investigate the two-way-coupled dynamics between the oscillatory flow in the
150 spinal SAS, represented by an infinite channel of constant thickness, and the oscillatory
151 flow in the syrinx, represented by a slender rectangular cavity, with an impermeable
152 elastic membrane subject to longitudinal tension used to model the thin layer of spinal-
153 cord tissue separating both spaces. As indicated in figure 2, this 2D configuration,
154 chosen here to maximize analytic simplification, can be envisioned as an approximate
155 representation of extracanalicular syringes, with the rigid wall opposing the membrane
156 representing the internal spinal-cord tissue. It is worth mentioning that the use of the
157 2D model neglects the hoop stresses induced by the azimuthal stretching of the tube,
158 which can be important, especially for canalicular syringes, for which an axisymmetric
159 configuration appears to be a more appropriate model. Also note that, by using an
160 impermeable membrane, our analysis also neglects effects of transmedullary interstitial
161 flow (Støverud *et al.* 2016; Wei *et al.* 2017; Cardillo & Camporeale 2021), a reasonably
162 valid approximation in investigating the cavity sloshing flow, since its characteristic time
163 is much smaller than that associated with the slow interstitial velocities.

164 As shown below, simplifications afforded by the disparity of scales present in the prob-
165 lem enable a rigorous asymptotic treatment of the canonical configuration represented
166 in figure 2(c), leading to closed-form expressions for all quantities of interest. Although
167 the predictive capability of the model is limited by the degree of simplification, the
168 analysis provides insights into the oscillatory cavity motion, yielding results in qualitative
169 agreement with previous *in vivo* observations pertaining to the prevailing cavity-flow
170 frequency (Vinje *et al.* 2018). Our analytic approach enables a complete parametric
171 description of the resulting transmural pressure to be made, including influences of
172 cavity size and SAS-flow frequency, which can be instrumental in guiding future FSI
173 investigations addressing anatomically correct systems.

174 The rest of the paper is organized as follows. The mathematical formulation of the
175 problem and associated dimensionless governing parameters are presented in § 2. The os-
176 cillatory motion arising at leading order in the limit of small stroke lengths is investigated
177 in § 3. The closed-form expressions obtained are used to explore parametric dependences
178 of the sloshing motion. The analysis is extended to investigate non-sinusoidal flow rates,
179 as those found in the spinal canal. The steady motion arising at the following order in the
180 asymptotic description is presented in § 4. Expressions are obtained for the slow time-
181 averaged Lagrangian motion of the fluid, involving the sum of the cycle-averaged Eulerian
182 velocity and the Stokes drift, and also for the stationary transmembrane pressure,
183 representative of the transmural pressure difference investigated in previous numerical
184 studies (Bertram 2010; Heil & Bertram 2016; Bertram & Heil 2017). Finally, concluding
185 remarks are provided in § 5.

2. Formulation of the problem

2.1. Preliminary considerations

As a simplified model of the SAS/cavity configuration, let us consider a two-dimensional channel of width h_o separated from a cavity of width h_c and length $L \gg h_o \sim h_c$ by an elastic membrane, as sketched in figure 1(e). Both regions are filled with the same incompressible viscous fluid of density ρ and kinematic viscosity ν (for CSF, $\rho \simeq 10^3 \text{ kg/m}^3$ and $\nu \simeq 0.7 \times 10^{-6} \text{ m}^2/\text{s}$). The fluid moves along the channel with a prescribed flow rate that varies harmonically with time t' according to $Q'_o \cos(\omega t')$, with the motion featuring characteristic longitudinal velocities $u_c = Q'_o/h_o$ and order-unity values of the associated Womersley number

$$\alpha = (h_o^2 \omega / \nu)^{1/2}, \quad (2.1)$$

where ω denotes the angular frequency. The longitudinal pressure variations associated with the flow in the channel, of order $\rho u_c \omega L$ as follows from a balance between local acceleration and pressure gradient, induce membrane deformations that drive an oscillatory motion in the cavity. The analysis below addresses the distinguished limit in which there exists two-way coupling between the cavity motion and the departures from Womersley flow emerging in the channel.

The deformation of the membrane is to be characterized in terms of the local distance h to the rigid channel wall (see figure 1(e)). Its response to the transmembrane pressure difference will be described with the simple linear elastic equation $T \partial^2 h / \partial x'^2 = \Delta p'$, where T is the constant longitudinal tension, x' is the streamwise coordinate, and $\Delta p'$ is the pressure difference across the membrane induced by the fluid motion, with $\Delta p' = 0$ for $Q'_o = 0$, so that in the absence of motion the membrane remains flat (i.e. $h = h_o$). Volume conservation in the closed cavity implies that

$$\int_0^L (h_o - h) dx' = 0, \quad (2.2)$$

at any instant of time.

In the analysis, it is assumed that the characteristic stroke length of the oscillatory motion in the canal u_c/ω is much smaller than the cavity length L , so that their ratio

$$\varepsilon = u_c / (L\omega) \ll 1 \quad (2.3)$$

defines a small asymptotic parameter measuring the effects of convective acceleration (i.e. ε is the inverse of the relevant Strouhal number). The distinguished limit considered here involves values of the membrane tension of order $T \sim \rho \omega^2 L^4 / h_o$, for which the magnitude of the relative membrane deformation

$$\frac{h_o - h}{h_o} \sim \frac{\rho u_c \omega L^3}{T h_o}, \quad (2.4)$$

deduced from an order-of-magnitude analysis of the membrane elastic equation with $\Delta p' \sim \rho u_c \omega L$, is of order $(h_o - h)/h_o \sim \varepsilon$. The problem will be described with use of cartesian coordinates with longitudinal and transverse components (x, y) scaled with L and h_o , respectively, and accompanying velocity components (u, v) scaled with u_c and $u_c h_o / L$, the latter scaling following from continuity. The pressure variations will be scaled with $\rho u_c \omega L$ to give the variable p and the membrane displacement will be written in the dimensionless form $\xi = (h_o - h) / (\varepsilon h_o) \sim 1$. The superscripts o and c will be used to denote the values of u , v and p in the channel and in the cavity, respectively.

2.2. Dimensionless equations

224

225 In the slender-flow approximation, which applies with small relative errors of order
 226 $(h_o/L)^2$, viscous stresses associated with longitudinal velocity derivatives can be ne-
 227 glected in the first approximation along with transverse pressure differences, so that
 228 $p = p(x, t)$ with $t = \omega t'$. The problem reduces to the integration of

$$\frac{\partial u}{\partial x} + \frac{\partial v}{\partial y} = 0 \quad \text{and} \quad \frac{\partial u}{\partial t} + \varepsilon \left(u \frac{\partial u}{\partial x} + v \frac{\partial u}{\partial y} \right) = -\frac{\partial p}{\partial x} + \frac{1}{\alpha^2} \frac{\partial^2 u}{\partial y^2} \quad (2.5)$$

229

for $0 \leq x \leq 1$ with boundary conditions at the lateral boundaries

$$u^o = v^o = 0 \text{ at } y = 1 \quad \text{and} \quad u^o = v^o - \partial \xi / \partial t = 0 \text{ at } y = \varepsilon \xi \quad (2.6)$$

230

for the channel flow and

$$u^c = v^c = 0 \text{ at } y = -H \quad \text{and} \quad u^c = v^c - \partial \xi / \partial t = 0 \text{ at } y = \varepsilon \xi \quad (2.7)$$

231

for the cavity flow, where $H = h_c/h_o$ denotes the dimensionless cavity width.

232

233

Since the flow rate takes the prescribed value $\int_0^1 u^o dy = \cos t$ upstream and down-
 stream from the cavity, the velocity in the channel for $x < 0$ and $x > 1$ reduces to the
 234 familiar Womersley solution

$$u^o = \text{Re} \left\{ \frac{1 - \cosh[\alpha \sqrt{i}(y - 1/2)] / \cosh(\alpha \sqrt{i}/2)}{1 - \tanh(\alpha \sqrt{i}/2) / (\alpha \sqrt{i}/2)} e^{it} \right\} \quad \text{with} \quad v^o = 0. \quad (2.8)$$

235

236

237

For $0 < x < 1$, the local flow rate $Q^o(x, t) = \int_{\varepsilon \xi}^1 u^o dy$, different in general from the
 prescribed boundary value $Q^o = \cos t$, is related to the flow rate in the cavity $Q^c =$
 $\int_{-H}^{\varepsilon \xi} u^c dy$ by

$$\frac{\partial}{\partial x} \left(\int_{\varepsilon \xi}^1 u^o dy \right) = -\frac{\partial}{\partial x} \left(\int_{-H}^{\varepsilon \xi} u^c dy \right) = \frac{\partial \xi}{\partial t}, \quad (2.9)$$

238

obtained by integrating the first equation in (2.5). Using the known boundary values

$$Q^o = \int_{\varepsilon \xi}^1 u^o dy = \cos t \quad \text{and} \quad Q^c = \int_{-H}^{\varepsilon \xi} u^c dy = 0 \quad \text{at} \quad x = 0, 1 \quad (2.10)$$

239

in integrating (2.9) yields

$$\int_{-H}^{\varepsilon \xi} u^c dy = \cos t - \int_{\varepsilon \xi}^1 u^o dy = -\int_0^x \frac{\partial \xi}{\partial t} d\hat{x}, \quad (2.11)$$

240

241

242

where \hat{x} represents a dummy integration variable. The above expression reveals that the
 flow rate in the cavity is balanced by a reverse flow in the channel of the same magnitude,
 so that the sum of both remains equal to the Womersley value $\cos t$.

243

The cavity and channel motions are coupled through the elastic equation

$$\mathcal{T} \frac{\partial^2 \xi}{\partial x^2} = p^o - p^c, \quad (2.12)$$

244

with the membrane deformation ξ satisfying the boundary conditions

$$\xi = 0 \quad \text{at} \quad x = 0, 1 \quad (2.13)$$

245

along with the integral constraint

$$\int_0^1 \xi dx = 0, \quad (2.14)$$

246 consistent with (2.2). In the elastic equation, the factor

$$\mathcal{T} = \frac{Th_o}{\rho\omega^2L^4} \quad (2.15)$$

247 is a dimensionless membrane tension.

2.3. Governing parameters and solution procedure

249 Besides the geometrical parameter $H = h_c/h_o$, the problem formulated above displays
 250 three parameters, namely, the Womersley number α defined in (2.1), the dimensionless
 251 stroke length ε defined in (2.3), and the dimensionless membrane tension \mathcal{T} defined
 252 in (2.15). The canonical model is designed to represent the dynamical behavior encoun-
 253 tered in syringomyelia syrinxes, with transverse sizes h_c comparable to, or somewhat
 254 larger than, the thickness of the surrounding SAS $h_o \sim 1 - 4$ mm, so that the focus
 255 below will be on order-unity values of H . For cardiac-driven flow, the angular frequency
 256 is of order $\omega = 2\pi \text{ s}^{-1}$ (i.e. assuming a cardiac rate of 60 beats per minute), so that
 257 the resulting Womersley number typically lies in the range $3 \lesssim \alpha \lesssim 12$, as follows
 258 from (2.1) when the value $\nu \simeq 0.7 \times 10^{-6} \text{ m}^2/\text{s}$ of the CSF kinematic viscosity at
 259 normal body temperature is used in the evaluation. With CSF peak velocities on the
 260 order of a few cm per second in the cervical SAS and cavity lengths of the order
 261 of a few cm, the resulting stroke length $\varepsilon = u_c/(\omega L)$ is moderately small (i.e. $\varepsilon \simeq$
 262 $0.1 - 0.2$), motivating an asymptotic description leveraging the limit $\varepsilon \ll 1$. The value
 263 of the dimensionless membrane tension \mathcal{T} must be selected to represent the dynamical
 264 deformation of the spinal cord tissue. The previous *in vivo* measurements of Vinje *et al.*
 265 (2018) reveal velocities in the syrinx that are comparable to those in the SAS, which in
 266 our model problem require membrane displacements ξ of order unity (e.g. see (2.11)) and
 267 corresponding values of \mathcal{T} also of order unity, according to (2.12). It appears therefore
 268 reasonable to explore the distinguished limit $\mathcal{T} \sim 1$ in which the channel and cavity flows
 269 display two-way coupling. Note that this limit arises when the characteristic wavelength
 270 $\lambda_e = [(Th_o)/(\rho\omega^2)]^{1/4}$ of the elastic membrane deformations associated with a forcing
 271 frequency ω is comparable to the cavity length L .

272 In the following quantitative description, pertaining to general order-unity values of H ,
 273 α , and \mathcal{T} and asymptotically small values of ε , all dependent variables will be expressed
 274 as expansions in powers of $\varepsilon \ll 1$ (e.g. $u^o = u_0^o + \varepsilon u_1^o + \dots$), leading to a hierarchy
 275 of problems that can be solved sequentially. The leading-order terms in the expansions,
 276 satisfying a linear problem, are purely harmonic, so that their cycle-averaged values
 277 are identically zero. In contrast, the first-order velocity corrections contain a non-zero
 278 steady-streaming component involving a non-zero transmembrane pressure difference, to
 279 be determined below. To facilitate the development, it is convenient to replace y with a
 280 normalized transverse coordinate η defined as

$$\eta = \frac{y - \varepsilon\xi}{1 - \varepsilon\xi} \quad (\text{channel}) \quad \text{and} \quad \eta = -\frac{y - \varepsilon\xi}{H + \varepsilon\xi} \quad (\text{cavity}), \quad (2.16)$$

281 such that $\eta = 0$ at the membrane and $\eta = 1$ at the opposite flat wall.

3. Leading-order oscillatory motion

3.1. Velocity field

284 The leading-order solution can be expressed in the form

$$(u_0^o, v_0^o, p_0^o, u_0^c, v_0^c, p_0^c, \xi_0) = \text{Re}[(U, V, P, \tilde{U}, \tilde{V}, \tilde{P}, \chi)e^{it}] \quad (3.1)$$

285 in terms of the complex functions $U(x, \eta)$, $V(x, \eta)$, $P(x)$, $\tilde{U}(x, \eta)$, $\tilde{V}(x, \eta)$, $\tilde{P}(x)$, and
 286 $\chi(x)$. In the channel, the solution reduces to the integration of

$$\frac{\partial U}{\partial x} + \frac{\partial V}{\partial \eta} = 0 \quad \text{and} \quad \frac{1}{\alpha^2} \frac{\partial^2 U}{\partial \eta^2} - iU = \frac{dP}{dx}, \quad (3.2)$$

287 with boundary conditions $U = V = 0$ at $\eta = 1$, $U = V - i\chi = 0$ at $\eta = 0$, as follows at this
 288 order from (2.5) and (2.6), with the reduced velocity satisfying the additional constraint
 289 $\int_0^1 U d\eta = 1$ at $x = 0, 1$, consistent with (2.10). Integrating the second equation in (3.2)
 290 with $U = 0$ at $\eta = (0, 1)$ yields

$$U = i \left\{ 1 - \frac{\cosh[\Lambda(2\eta - 1)]}{\cosh \Lambda} \right\} \frac{dP}{dx} \quad (3.3)$$

291 where $\Lambda = \alpha\sqrt{i}/2$. The expression for U can be used in the first equation in (3.2) to
 292 provide

$$V = -i \left\{ \eta - 1 - \frac{\sinh[\Lambda(2\eta - 1)] - \sinh \Lambda}{2\Lambda \cosh \Lambda} \right\} \frac{d^2 P}{dx^2} \quad (3.4)$$

293 upon integration with use of $V = 0$ at $\eta = 1$.

The same integration procedure can be applied to the cavity flow to give

$$\tilde{U} = i \left\{ 1 - \frac{\cosh[H\Lambda(2\eta - 1)]}{\cosh(H\Lambda)} \right\} \frac{d\tilde{P}}{dx}, \quad (3.5)$$

$$\tilde{V} = iH \left\{ \eta - 1 - \frac{\sinh[H\Lambda(2\eta - 1)] - \sinh(H\Lambda)}{2H\Lambda \cosh(H\Lambda)} \right\} \frac{d^2 \tilde{P}}{dx^2}. \quad (3.6)$$

294 The velocity profiles (3.3) and (3.5) can be used to evaluate the integrals

$$\int_0^1 U d\eta = \frac{1}{\beta} \frac{dP}{dx} \quad \text{and} \quad H \int_0^1 \tilde{U} d\eta = \frac{1}{\tilde{\beta}} \frac{d\tilde{P}}{dx}, \quad (3.7)$$

295 which enter in the computation of the leading-order oscillatory flow rates

$$Q_0^o = \int_0^1 u_0^o dy = \text{Re} \left(\int_0^1 U d\eta e^{it} \right) \quad \text{and} \quad Q_0^c = \int_{-H}^0 u_0^c dy = \text{Re} \left(H \int_0^1 \tilde{U} d\eta e^{it} \right), \quad (3.8)$$

296 with

$$\beta = -i [1 - \Lambda^{-1} \tanh \Lambda]^{-1} \quad \text{and} \quad \tilde{\beta} = -i [H - \Lambda^{-1} \tanh(H\Lambda)]^{-1}. \quad (3.9)$$

297 As can be seen from (3.3) and (3.4), when the pressure gradient takes the uniform
 298 unperturbed value $dP/dx = \beta$, the leading-order velocity in the channel (u_0^o, v_0^o) =
 299 $\text{Re}[(U, V)e^{it}]$ reduces to the familiar Womersley solution (2.8) existing for $x < 0$ and
 300 $x > 1$.

3.2. Membrane deformation

301 The pressure distributions in the channel and in the cavity $P(x)$ and $\tilde{P}(x)$, which
 302 complete the determination of the flow at this order, are related to the membrane
 303 deformation by
 304

$$\frac{d^2 P}{dx^2} = i\beta\chi \quad \text{and} \quad \frac{d^2 \tilde{P}}{dx^2} = -i\tilde{\beta}\chi, \quad (3.10)$$

305 as follows from using the boundary conditions $V = \tilde{V} = i\chi$ at $\eta = 0$ in (3.4) and (3.6).
 306 Their values are coupled through

$$\mathcal{T} \frac{d^2 \chi}{dx^2} = P - \tilde{P}, \quad (3.11)$$

307 obtained at leading-order from (2.12). Differentiating twice the above equation followed
 308 by substitution of (3.10) provides the boundary-value problem

$$\frac{d^4 \chi}{dx^4} - \frac{i(\beta + \tilde{\beta})}{\mathcal{T}} \chi = 0 \quad \text{with} \quad \begin{cases} d^3 \chi / dx^3 = \beta / \mathcal{T} \\ \chi = 0 \end{cases} \quad \text{at} \quad x = (0, 1) \quad (3.12)$$

309 for the membrane displacement χ . The boundary condition involving the third derivative
 310 follows from imposing the conditions $dP/dx - \beta = d\tilde{P}/dx = 0$ at $x = 0, 1$, corresponding
 311 to $\int_0^1 U d\eta - 1 = \int_0^1 \tilde{U} d\eta = 0$. The deformation satisfies $\int_0^1 \chi dx = 0$, as can be readily
 312 verified by performing a first quadrature of (3.12).

313 The solution to (3.12) can be written as

$$\chi = \frac{\beta}{\mathcal{T}} \left\{ \frac{\sin\left(\frac{\gamma}{2\mathcal{T}^{1/4}}\right) \sinh\left[\frac{\gamma}{\mathcal{T}^{1/4}}\left(x - \frac{1}{2}\right)\right] - \sinh\left(\frac{\gamma}{2\mathcal{T}^{1/4}}\right) \sin\left[\frac{\gamma}{\mathcal{T}^{1/4}}\left(x - \frac{1}{2}\right)\right]}{\left(\frac{\gamma}{\mathcal{T}^{1/4}}\right)^3 \left[\sinh\left(\frac{\gamma}{2\mathcal{T}^{1/4}}\right) \cos\left(\frac{\gamma}{2\mathcal{T}^{1/4}}\right) + \cosh\left(\frac{\gamma}{2\mathcal{T}^{1/4}}\right) \sin\left(\frac{\gamma}{2\mathcal{T}^{1/4}}\right)\right]} \right\}, \quad (3.13)$$

314 where $\gamma = [i(\beta + \tilde{\beta})]^{1/4}$. The above expression can be used in (3.10) to obtain $d^2 P/dx^2$
 315 and $d^2 \tilde{P}/dx^2$, needed in (3.4) and (3.6). On the other hand, integration of (3.10) subject
 316 to $dP/dx - \beta = d\tilde{P}/dx = 0$ at $x = 0$ provides the pressure gradients required in (3.3)
 317 and (3.5), resulting in

$$\frac{1}{\beta} \frac{dP}{dx} - 1 = -\frac{1}{\tilde{\beta}} \frac{d\tilde{P}}{dx} = i \int_0^x \chi d\hat{x} \quad (3.14)$$

with

$$i \int_0^x \chi d\hat{x} = \frac{\beta}{\beta + \tilde{\beta}} \left\{ \coth[\gamma/(2\mathcal{T}^{1/4})] + \cot[\gamma/(2\mathcal{T}^{1/4})] \right\}^{-1} \\ \times \left(\frac{\cosh\left[\frac{\gamma}{\mathcal{T}^{1/4}}\left(x - \frac{1}{2}\right)\right] - \cosh\left(\frac{\gamma}{2\mathcal{T}^{1/4}}\right)}{\sinh[\gamma/(2\mathcal{T}^{1/4})]} + \frac{\cos\left[\frac{\gamma}{\mathcal{T}^{1/4}}\left(x - \frac{1}{2}\right)\right] - \cos\left(\frac{\gamma}{2\mathcal{T}^{1/4}}\right)}{\sin[\gamma/(2\mathcal{T}^{1/4})]} \right), \quad (3.15)$$

318 the latter entering when using (3.7) and (3.8) for the determination of the flow rates

$$\int_{-H}^0 u_0^c dy = \cos t - \int_0^1 u_0^o dy = -\text{Re} \left(i \int_0^x \chi d\hat{x} e^{it} \right). \quad (3.16)$$

319 Note that the last equation corresponds to the leading-order form of (2.11).

3.3. Oscillatory motion

321 The closed-form expressions derived above can be used to investigate the main features
 322 of the FSI oscillatory dynamics and its parametric dependences. We begin by plotting in
 323 the middle and right-hand-side panels of figure 3 snapshots of streamlines and membrane
 324 displacement at two different instants of time corresponding to a configuration with $\alpha = 5$
 325 and $H = 1$. Color contours are used to represent the associated vorticity, which in the
 326 slender-flow approximation reduces to $-\partial u_0 / \partial y$. The accompanying temporal variation
 327 of the leading-order flow rates $Q_0^c = \int_{-H}^0 u_0^c dy$ and $Q_0^o = \int_0^1 u_0^o dy$ at the canal middle
 328 section $x = 0.5$ are shown in the left panels. The computations reveal, in particular,
 329 that the value of \mathcal{T} needs to be much smaller than unity to induce significant membrane
 330 displacements (and therefore significant motion in the cavity). For example, for $\mathcal{T} = 0.05$,

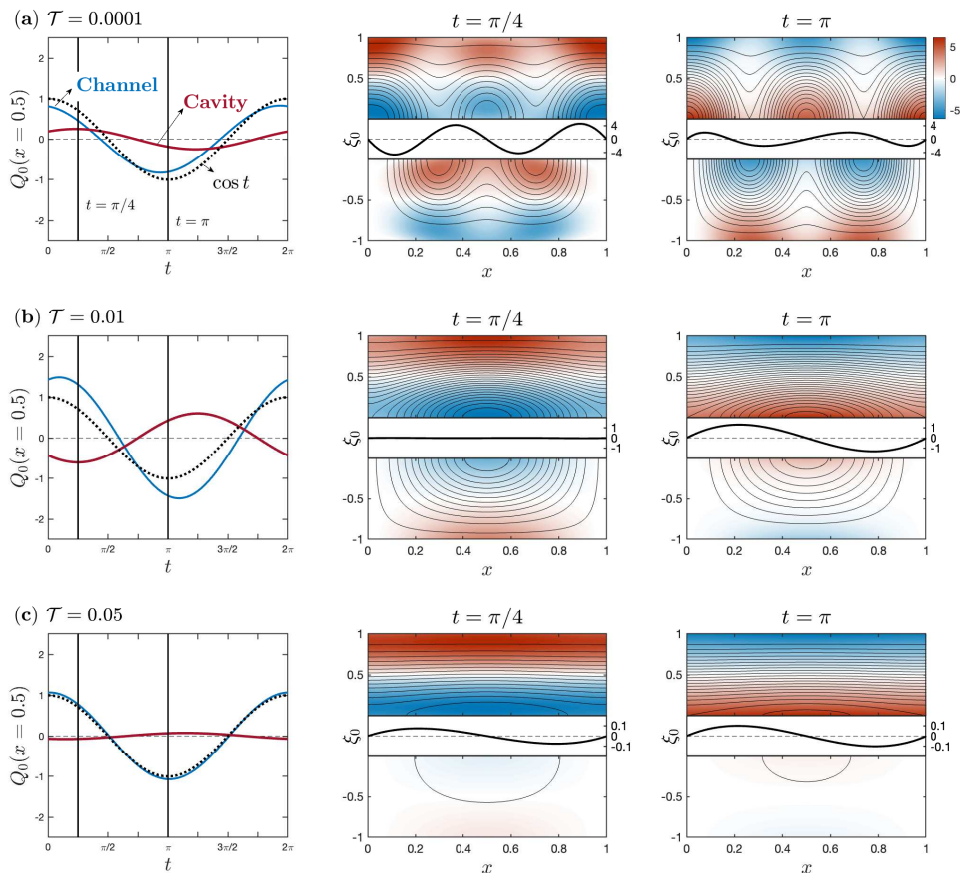


FIGURE 3. Oscillatory flow for a configuration with $\alpha = 5$, $H = 1$, and $\mathcal{T} =$ (a) 0.0001, (b) 0.01, and (c) 0.05. The left plots show the variation with time of the channel (blue) and cavity (red) flow rates $Q_0^c = \int_0^1 u_0^c dy$ and $Q_0^c = \int_{-H}^0 u_0^c dy$ at $x = 0.5$ evaluated using (3.16), while the middle and right plots show streamlines and color contours of vorticity at $t = \pi/4$ and $t = \pi$ along with the corresponding membrane displacement ξ_0 . To facilitate comparisons, a fixed constant streamline spacing of $\delta\psi_0 = 0.05$ has been used in representing the streamlines, with the stream function ψ_0 computed using $\partial\psi_0/\partial y = u_0$ and $\partial\psi_0/\partial x = -v_0$.

331 the case shown in figure 3(c), the membrane displacement is limited to values $\xi_0 < 0.1$ and
 332 the fluid remains nearly stagnant in the cavity, associated departures from Womersley
 333 flow in the channel being correspondingly small.

334 The limited membrane displacement found for $\mathcal{T} \sim 1$ can be attributed to the smallness
 335 of the term in curly brackets in the general expression (3.13). This can be seen more
 336 clearly by considering the limit of very stiff membranes $\mathcal{T} \gg 1$, in which one can readily
 337 integrate (3.12) to give the approximate result

$$\chi \simeq \frac{\beta}{\mathcal{T}} \frac{x}{6} \left(x - \frac{1}{2} \right) (x - 1) \quad \text{for } \mathcal{T} \gg 1. \quad (3.17)$$

338 Straightforward evaluation reveals that the maximum displacement in this limit, reached
 339 at $x = 1/2 \pm \sqrt{3}/6$, is $\chi \simeq 8.02 \times 10^{-3} \beta/\mathcal{T}$, with the small numerical factor being
 340 consistent with the results shown in the figure.

341 In contrast to the case $\mathcal{T} = 0.05$, the configurations with $\mathcal{T} = 10^{-4}$ and $\mathcal{T} = 10^{-2}$,

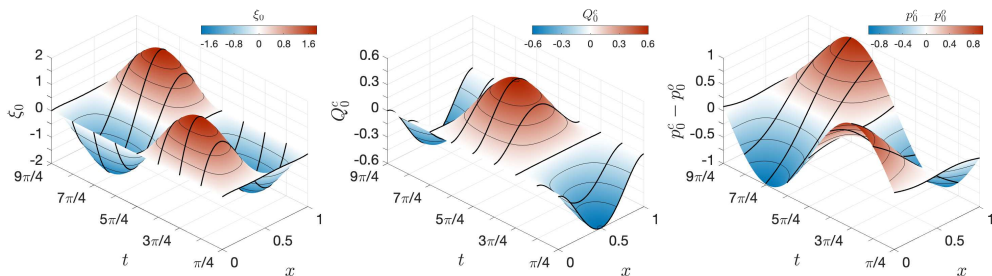


FIGURE 4. The variation with time of the membrane displacement ξ_0 , cavity flow rate $Q_0^c = \int_{-H}^0 u_0^c dy$, and oscillatory transmembrane pressure difference $p_0^c - p_0^o$ for a cavity with $\alpha = 5$, $H = 1$, and $\mathcal{T} = 0.01$.

shown in figures 3(a) and 3(b), respectively, show velocities in the cavity that are comparable to those in the channel. The streamlines in all plots have been represented using the same values of the stream function, so that their inter-spacing characterizes the local flow speed. The comparison of the streamlines in figures 3(a) and 3(b) reveals that the flow patterns become more complicated as the membrane becomes more flexible for decreasing values of \mathcal{T} . In interpreting this result, it is worth recalling that the dimensionless membrane tension can be expressed as $\mathcal{T} = (\lambda_e/L)^4$ in terms of the characteristic elastic wavelength $\lambda_e = [(Th_o)/(\rho\omega^2)]^{1/4}$, so that the number of membrane undulations increases for decreasing values of \mathcal{T} , driving separate regions of recirculating flow.

The dynamics of the sloshing motion induced in the cavity is characterized in figure 4 by plotting the temporal variation over a cycle of the tightly coupled cavity deformation ξ_0 , flow rate $Q_0^c = \int_{-H}^0 u_0^c dy$, and oscillatory transmembrane pressure $p_0^c - p_0^o = \text{Re}[(\tilde{P} - P)e^{it}]$, with $\tilde{P} - P$ evaluated from (3.11) by straightforward double differentiation of (3.13). As can be expected from (3.11) and (3.16), the membrane displacement is in phase with $p_0^c - p_0^o$, while the flow rate is in quadrature. At the initial time $t = \pi/4$ selected in the figure, the membrane is practically flat and the transmembrane pressure difference is very small. The fluid, with an initially negative flow rate, moves upstream, deforming the membrane and inducing a negative pressure gradient that slows down the motion, so that the velocity vanishes when the deformation reaches its maximum at $t = 3\pi/4$. The flow reverses for $t > 3\pi/4$, with the negative pressure gradient driving the flow downstream. A nearly flat membrane with negligible transmembrane pressure gradient is found for $t = 5\pi/4$ as the flow rate reaches its peak positive value. The sloshing behavior is replicated over the second half of the cycle following the expected sinusoidal pattern. In view of figure 3(a), it can be anticipated that the sloshing-flow structure becomes more complicated as the elastic wavelength becomes much smaller than L for decreasing values of \mathcal{T} , that being the case investigated below.

3.4. Very flexible membranes

For values of \mathcal{T} smaller than those considered in figure 3, the membrane undulations, of larger amplitude for decreasing \mathcal{T} , remain mostly confined to near-edge regions scaling with the elastic-wave wavelength. Illustrative results pertaining to this limit of very flexible membranes are shown in figure 5, including instantaneous membrane shapes at selected times and associated cavity flow rates.

The structure that emerges can be investigated by exploring the asymptotic limit $\mathcal{T} \ll 1$, wherein equation (3.12) reduces to $\chi = 0$ while (3.11) yields $P = \tilde{P}$, so that

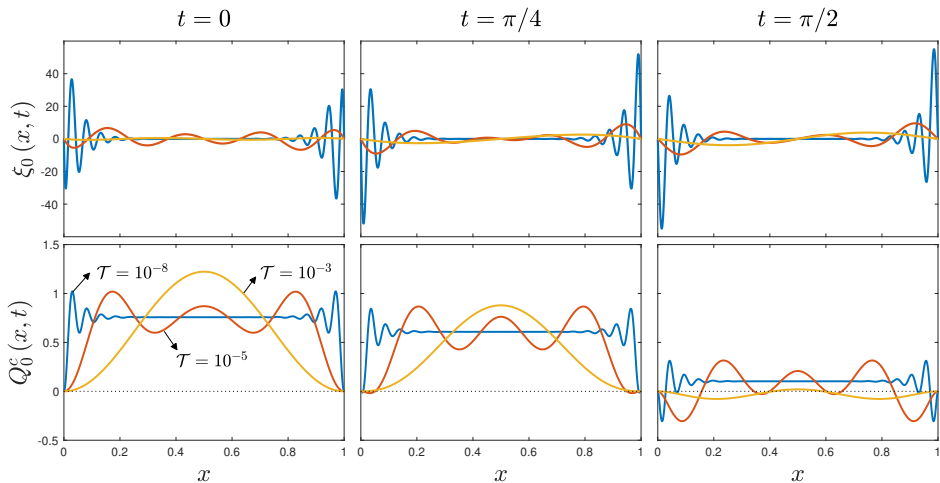


FIGURE 5. The streamwise variation of the membrane displacement ξ_0 and cavity flow rate $Q_0^c = \int_{-H}^0 u_0^c dy$ at $t = (0, \pi/4, \pi/2)$ for $\alpha = 3$, $H = 2$ and $\mathcal{T} = (10^{-3}, 10^{-5}, 10^{-8})$.

377 the fluid moves in the channel and in the cavity under the action of the same pressure
 378 gradient. This solution fails near the edges of the membrane, in two boundary regions $x \sim$
 379 $\mathcal{T}^{1/4} \ll 1$ and $(1-x) \sim \mathcal{T}^{1/4} \ll 1$ where $\chi \sim \mathcal{T}^{-1/4} \gg 1$ and $P \sim \tilde{P} \sim \mathcal{T}^{1/4} \ll 1$ whose
 380 solution determines the pressure gradient driving the uniform flow rate in the central
 381 region. Introducing the rescaled variables $\zeta = x/\mathcal{T}^{1/4}$ (replaced by $\zeta = (1-x)/\mathcal{T}^{1/4}$
 382 in the description of the right-hand-side edge region), $\chi_e = \mathcal{T}^{1/4}\chi$, $P_e = P/\chi^{1/4}$, and
 383 $\tilde{P}_e = \tilde{P}/\chi^{1/4}$ leads to the modified boundary-value problem

$$\frac{d^4 \chi_e}{d\zeta^4} - \gamma^4 \chi_e = 0 \quad \begin{cases} \frac{d^3 \chi_e}{d\zeta^3} - \beta = \chi_e = 0 & \text{at } \zeta = 0, \\ \chi_e \rightarrow 0 & \text{as } \zeta \rightarrow \infty, \end{cases} \quad (3.18)$$

384 which can be integrated to give

$$\chi_e = \frac{\beta}{\gamma^3(1-i)} (e^{i\gamma\zeta} - e^{-\gamma\zeta}). \quad (3.19)$$

385 Without loss of generality, in writing the above expression we have used the complex
 386 root $\gamma = [i(\beta + \tilde{\beta})]^{1/4}$ lying in the first quadrant, so that $e^{i\gamma\zeta} \rightarrow 0$ and $e^{-\gamma\zeta} \rightarrow 0$ as
 387 $\zeta \rightarrow \infty$. Substituting (3.19) into the rescaled form of (3.14) and (3.16) yields

$$\frac{1}{\beta} \frac{dP_e}{d\zeta} - 1 = -\frac{1}{\tilde{\beta}} \frac{d\tilde{P}_e}{d\zeta} = \frac{\beta}{\beta + \tilde{\beta}} \left(\frac{e^{-\gamma\zeta} - ie^{i\gamma\zeta}}{1-i} - 1 \right) \quad (3.20)$$

388 and

$$\int_{-H}^0 u_0^c dy = \cos t - \int_0^1 u_0^o dy = -\text{Re} \left[\frac{\beta}{\beta + \tilde{\beta}} \left(\frac{e^{-\gamma\zeta} - ie^{i\gamma\zeta}}{1-i} - 1 \right) e^{it} \right] \quad (3.21)$$

389 for the pressure gradients and flow rates in the near-edge regions. The result can be
 390 evaluated as $\zeta \rightarrow \infty$ to obtain the uniform values

$$\frac{dP}{dx} = \frac{d\tilde{P}}{dx} = \frac{\beta\tilde{\beta}}{\beta + \tilde{\beta}} \quad (3.22)$$

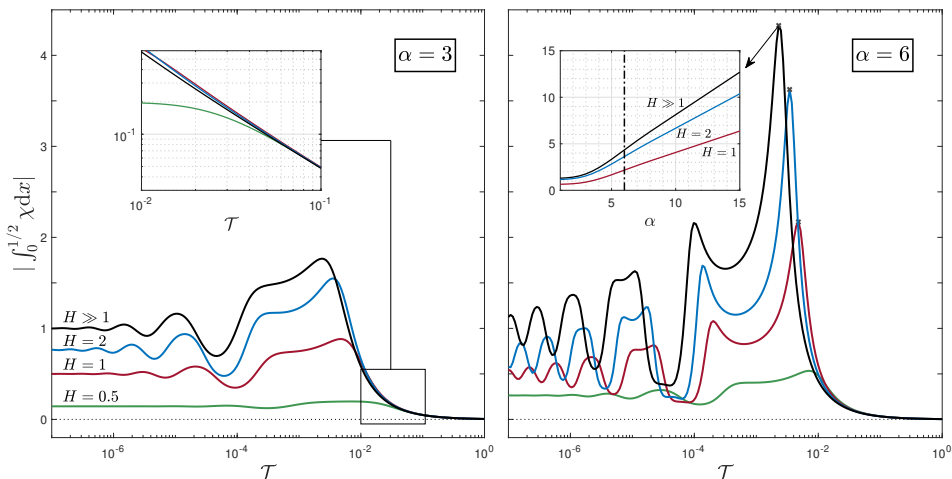


FIGURE 6. The variation with \mathcal{T} of the amplitude of the oscillating flow rate $|\int_0^{1/2} \chi dx|$ across the central section $x = 1/2$ of the cavity for $\alpha = 3$ (left) and $\alpha = 6$ (right) and four different values of $H = (0.5, 1, 2, \infty)$. The inset in the left panel represents an expanded view of the curves as they merge for increasing \mathcal{T} while that in the right panel gives the variation with α of the peak value of $|\int_0^{1/2} \chi dx|$ for three different values of H .

391 and

$$\int_{-H}^0 u_0^c dy = \cos t - \int_0^1 u_0^d dy = \operatorname{Re} \left(\frac{\beta e^{it}}{\beta + \tilde{\beta}} \right) \quad (3.23)$$

392 that prevail away from the edge regions.

393 3.5. Parametric dependences of the flow rate

394 As can be inferred from (3.16), the parametric dependences of the oscillating flow rate
 395 in the cavity (and, correspondingly, of the departures from Womersley flow in the channel)
 396 are embodied in the function $i \int_0^x \chi d\hat{x}$ given in (3.15). A measure of the induced motion
 397 is provided by the local amplitude of the oscillating flow rate across the central section
 398 $x = 1/2$ of the cavity, given by the modulus $|\int_0^{1/2} \chi dx|$, which is also proportional to the
 399 corresponding stroke volume $\int_t^{t+2\pi} |Q_0^c(1/2, t)| dt / (2\pi) = (2/\pi) |\int_0^{1/2} \chi dx|$. The variation
 400 of $|\int_0^{1/2} \chi dx|$ with \mathcal{T} is represented in figure 6 for different values of H and α .

401 The curves reproduce the trends previously identified. In particular, the motion is
 402 very limited for values of $\mathcal{T} \gtrsim 0.1$, when the flow rate becomes independent of H , as seen
 403 in the inset of the left-hand-side panel, with a value that decays for $\mathcal{T} \gg 1$ according
 404 to $|\int_0^{1/2} \chi dx| = |\beta| / (384\mathcal{T})$, a result derived with use of (3.17). In the opposite limit
 405 $\mathcal{T} \ll 1$ of very flexible membranes, the flow-rate amplitude approaches the constant
 406 value $|\int_0^{1/2} \chi dx| = |\beta| / (\beta + \tilde{\beta})$, larger for larger H , with $|\int_0^{1/2} \chi dx| = 0.5$ when $\beta = \tilde{\beta}$
 407 for $H = 1$. The flow rate in the central part of the cavity becomes maximum for an
 408 intermediate value of \mathcal{T} lying in the range $10^{-3} < \mathcal{T} < 10^{-2}$, with the peak becoming
 409 more pronounced with increasing α , as shown in the inset of the right-hand-side panel.
 410 Between their peak values and the asymptotic values approached as $\mathcal{T} \rightarrow 0$, the curves in
 411 figure 6 display oscillations of decreasing amplitude, which are related to the development
 412 of an increasing number of membrane undulations as the cavity length L becomes larger
 413 than the elastic wavelength λ_e for decreasing values of $\mathcal{T} = (\lambda_e/L)^4$.

3.6. Effects of complex waveform

414

415 The rapid decay from its peak value experienced by $|\int_0^{1/2} \chi dx|$ as \mathcal{T} increases, more
 416 prominent for larger α , is indicative of a strong frequency dependence of the flow rate
 417 induced in the cavity. As indicated by the plots in figure 6, for intermediate values of
 418 $\mathcal{T} \sim 10^{-2}$, increasing the frequency (i.e. reducing the value of $\mathcal{T} \propto \omega^{-2}$ and increasing the
 419 value of $\alpha \propto \omega^{1/2}$) may promote significantly the motion in the cavity, with implications
 420 concerning the characteristics of the oscillatory flow in syringomyelia syringes, an aspect
 421 of the flow investigated below.

422 The typical waveform of the cardiac-driven flow rate Q' at the entrance of the spinal
 423 canal has a non-sinusoidal waveform, so that the Fourier decomposition of the signal
 424 has multiple harmonics of frequency $n\omega$. For instance, a Fourier analysis of the periodic
 425 flow rate corresponding to a Chiari patient, shown in figure 7(a), obtained by rescaling
 426 phase-contrast (PC) MRI velocity measurements reported by Vinje *et al.* (2018), yields

$$Q'/\langle|Q'|\rangle = \sum_{n=1}^{\infty} \text{Re}(A_n e^{int}) \quad (3.24)$$

427 where A_n are complex constants of order unity, with $A_1 = 0.2765 - 1.4686i$, $A_2 =$
 428 $0.0206 - 0.6748i$ and $A_3 = -0.1203 - 0.2222i$ for the first three modes. Here, we
 429 have normalized the flow rate with its average amplitude $\langle|Q'|\rangle = \int_0^{2\pi} |Q'| dt / (2\pi)$. For
 430 comparison, figure 7(a) includes the purely sinusoidal case $Q'/\langle|Q'|\rangle = (\pi/2) \sin(t)$ (i.e.
 431 $A_1 = -(\pi/2)i$ with $A_n = 0$ for $n > 1$).

432 The analysis given above, pertaining to a simple sinusoidal flow rate, can be readily
 433 extended to account for the presence of the different harmonics, leading to the flow-rate
 434 expressions

$$\int_{-H}^0 u_c^o dy = \sum_{n=1}^{\infty} \text{Re}(A_n e^{int}) - \int_0^1 u_0^o dy = -\text{Re} \left(\sum_{n=1}^{\infty} A_n i \int_0^x \chi_n d\hat{x} e^{int} \right), \quad (3.25)$$

435 with $u_c = \langle|Q'|\rangle/h_o$ used as characteristic velocity in scaling the problem. The value
 436 of $i \int_0^x \chi_n d\hat{x}$, measuring the amplification of a specific mode n , can be determined from
 437 the general expression (3.15) by simply replacing \mathcal{T} with \mathcal{T}/n^2 and evaluating β , $\tilde{\beta}$, and
 438 $\gamma = [i(\beta + \tilde{\beta})]^{1/4}$ with use of $n^{1/2}\alpha$ in place of α .

439 Bearing in mind the frequency dependence discussed above in connection with figure 6,
 440 one may anticipate that, for configurations with \mathcal{T} sufficiently large, higher-order har-
 441 monics $n > 1$ may have values of the amplification factor $i \int_0^x \chi_n d\hat{x}$ that are larger than
 442 those of the fundamental frequency, that being a result of the variation of the frequency-
 443 weighted membrane tension \mathcal{T}/n^2 and Womersley number $n^{1/2}\alpha$. As a consequence,
 444 although the fundamental mode with frequency ω is clearly dominant in the flow rate at
 445 the entrance of the spinal canal Q' , so that the waveform is nearly sinusoidal, as shown
 446 in figure 7(a), the motion induced in the syrinx may exhibit pronounced oscillations
 447 at higher frequencies $n\omega$. As previously discussed in the introduction, such dynamics
 448 has been observed in *in vivo* non-invasive measurements performed in syringomyelia
 449 patients both before and after craniovertebral decompression (Vinje *et al.* 2018). In the
 450 preoperative study, the flow in the syrinx was found to display three full oscillations per
 451 cardiac cycle (i.e. Vinje *et al.* (2018) report 210 cycles per minute for a heart rate of 73
 452 beats per minute), indicating that the third harmonic $n = 3$ was dominant. In contrast,
 453 two months after surgery, the flow in the syrinx, now reduced in size (i.e. corresponding
 454 to a smaller value of H in our analysis), exhibited instead two full oscillations per cardiac

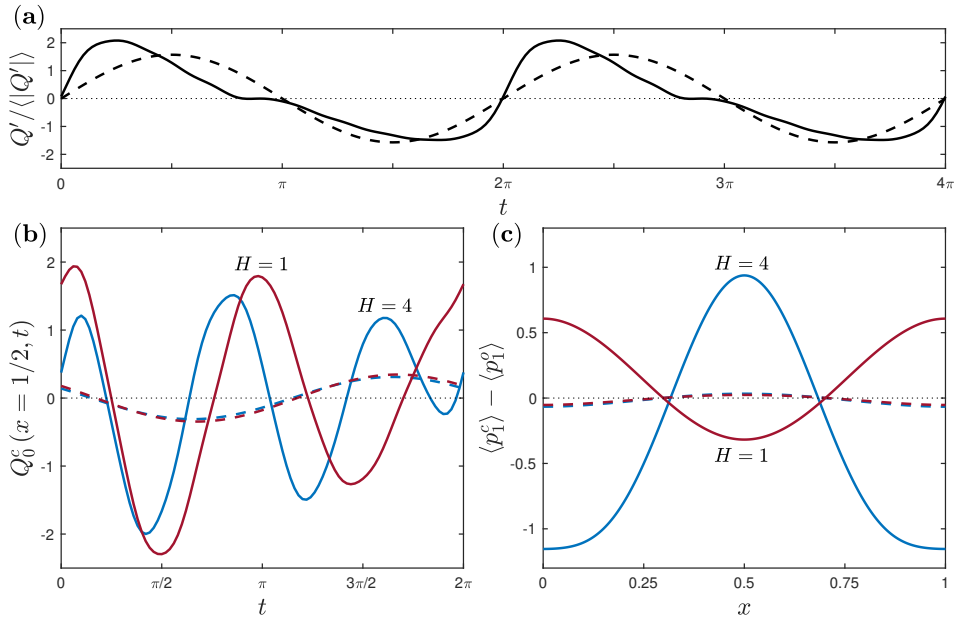


FIGURE 7. The upper panel (a) compares the dimensionless flow rate at the entrance of the spinal canal measured by cardiac-gated PC MRI (adapted from Vinje *et al.* (2018)) (solid curve) with the sinusoidal signal $Q'/|Q'| = (\pi/2) \sin(t)$ (dashed curve). The two wave forms are used to determine the response of the cavity flow for a configuration with $\alpha = 5$, $\mathcal{T} = 0.02$ and two different cavity widths $H = 1$ (red curves) and $H = 4$ (blue curves), including (b) the variation with time of the cavity flow rate $Q_0^c = \int_{-H}^0 u_0^c dy$ at $x = 1/2$ determined from (3.25) and (c) the streamwise variation of the transmurial steady pressure difference $\langle p_1^c \rangle - \langle p_1^o \rangle$ computed from (4.29). For consistency with (a), the solid/dashed curves in (b) and (c) are computed with the complex/sinusoidal channel-flow wave forms.

455 cycle (i.e. 200 cycles per minute for a heart rate of 97 beats per minute), consistent with
 456 the second harmonic $n = 2$ being dominant instead in the postoperative state.

457 The results of the simple FSI model developed here can be used to investigate this
 458 intriguing behavior. Results of an illustrative computation are given in figure 7 for a
 459 configuration with $\alpha = 5$, $\mathcal{T} = 0.02$ and two different values of H . Figure 7(b) shows
 460 the waveform of the periodic flow rate $Q_0^c = \int_0^1 u_0^c dy$ across the cavity middle section
 461 $x = 1/2$ as determined from (3.25) using the sinusoidal flow rate $Q'/|Q'| = (\pi/2) \sin(t)$
 462 (dashed curves) and using ten modes in the Fourier expansion (3.24) for the spinal-canal
 463 flow rate represented with the solid curve in figure 7(a) (solid curves). As can be seen,
 464 the flow rate induced in the cavity when the channel flow is purely sinusoidal follows
 465 the fundamental frequency. In contrast, the cavity-flow response to the complex wave
 466 form, of much larger amplitude, exhibits multiple cycles. In particular, it is seen that
 467 the curve with $H = 4$, representative of the preoperative state, exhibits three cycles,
 468 in agreement with the previous *in vivo* observations (Vinje *et al.* 2018). Interestingly,
 469 when the width of the cavity is reduced to $H = 1$, mimicking the reduction in syringe
 470 transverse size that proceeds surgery, the second harmonic becomes dominant, so that the
 471 resulting waveform of the cavity flow rate shows two cycles instead, again in agreement
 472 with the observations (Vinje *et al.* 2018). It is remarkable that, while the configuration
 473 investigated here is much too simple to enable quantitative predictions to be made, it is
 474 still able to reproduce some aspects of the observed *in vivo* dynamics when the value of
 475 \mathcal{T} is selected in the appropriate range.

4. Secondary motion

4.1. Steady streaming

The leading-order solution investigated in the preceding section has a zero time average, so that it does not result in a net transmembrane pressure difference. In contrast, the first-order corrections include a steady component, which can be determined by taking the time average of the corresponding governing equations, obtained by collecting terms of order ε in (2.5). In the channel, the problem reduces to the integration of

$$\frac{\partial \langle u_1^o \rangle}{\partial x} + \frac{\partial \langle v_1^o \rangle}{\partial \eta} + G(x, \eta) = 0 \quad \text{and} \quad \frac{1}{\alpha^2} \frac{\partial^2 \langle u_1^o \rangle}{\partial \eta^2} = \frac{d \langle p_1^o \rangle}{dx} + F(x, \eta) \quad (4.1)$$

subject to $\langle u_1^o \rangle = \langle v_1^o \rangle = 0$ at $\eta = 0, 1$ and $\int_0^1 \langle u_1^o \rangle d\eta = 0$ at $x = 0, 1$, with $\langle \cdot \rangle = \int_t^{t+2\pi} \cdot dt / (2\pi)$ representing the time-averaging operator. The steady motion is driven by the effect of convective acceleration and the nonlinear interactions stemming from the deformation of the canal, which enter in the problem through the functions

$$G = (\eta - 1) \left\langle \frac{\partial \xi_0}{\partial x} \frac{\partial u_0^o}{\partial \eta} \right\rangle + \left\langle \xi_0 \frac{\partial v_0^o}{\partial \eta} \right\rangle \quad \text{and} \quad (4.2)$$

$$F = (\eta - 1) \left\langle \frac{\partial \xi_0}{\partial t} \frac{\partial u_0^o}{\partial \eta} \right\rangle + \left\langle u_0^o \frac{\partial u_0^o}{\partial x} \right\rangle + \left\langle v_0^o \frac{\partial u_0^o}{\partial \eta} \right\rangle - \frac{2}{\alpha^2} \left\langle \xi_0 \frac{\partial^2 u_0^o}{\partial \eta^2} \right\rangle. \quad (4.3)$$

The time averages of products of harmonic functions in the above expressions can be written in terms of U , V , and χ with use of the identity $\langle \text{Re}(e^{it}A) \text{Re}(e^{it}B) \rangle = \text{Re}(AB^*)/2$, which applies to any generic time-independent complex functions A and B , with the asterisk $*$ denoting complex conjugates.

Because of the canal deformation, the cycle-averaged velocity is non-solenoidal, as seen in the first equation of (4.1), resulting in a nonzero flow rate $\langle Q_1^o \rangle = \int_0^1 \langle u_1^o \rangle d\eta$. Its value can be determined directly by integrating the continuity equation across the canal with $\langle v_1^o \rangle = 0$ at $\eta = 0, 1$ to give

$$\frac{d}{dx} \left[\int_0^1 \langle u_1^o \rangle d\eta - \left\langle \xi_0 \int_0^1 u_0^o d\eta \right\rangle \right] = 0 \quad (4.4)$$

after use is made of integration by parts to reduce $\int_0^1 G d\eta$. Since $\int_0^1 \langle u_1^o \rangle d\eta = 0$ at $x = 0, 1$, where $\xi_0 = 0$, it follows that

$$\int_0^1 \langle u_1^o \rangle d\eta = \left\langle \xi_0 \int_0^1 u_0^o d\eta \right\rangle. \quad (4.5)$$

As seen later in § 4.2, this non-zero flow rate is balanced exactly by that of the Stokes drift, so that the mean Lagrangian motion has a zero flow rate, as it should.

The steady-streaming velocity in the channel is computed by integrating the second equation in (4.1) subject to $\langle u_1^o \rangle = 0$ at $\eta = 0, 1$ to give

$$\langle u_1^o \rangle = -\alpha^2 \left[\frac{\eta}{2} (1 - \eta) \frac{d \langle p_1^o \rangle}{dx} + \int_0^\eta \hat{\eta} F d\hat{\eta} + \eta \left(\int_\eta^1 F d\hat{\eta} - \int_0^1 \eta F d\eta \right) \right], \quad (4.6)$$

which can be used in integrating the first equation in (4.1) with the condition $\langle v_1^o \rangle = 0$

at $\eta = 0$ to obtain

$$\begin{aligned} \langle v_1^o \rangle = \alpha^2 \frac{\partial}{\partial x} \left[\frac{\eta^2}{2} \left(\frac{1}{2} - \frac{\eta}{3} \right) \frac{d\langle p_1^o \rangle}{dx} - \frac{1}{2} \int_0^\eta F \hat{\eta}^2 d\hat{\eta} \right. \\ \left. + \eta \int_0^\eta F \hat{\eta} d\hat{\eta} + \frac{\eta^2}{2} \left(\int_\eta^1 F d\hat{\eta} - \int_0^1 F \eta d\eta \right) \right] - \int_0^\eta G d\hat{\eta}, \end{aligned} \quad (4.7)$$

497 where the pressure gradient is given by

$$\frac{d\langle p_1^o \rangle}{dx} = -\frac{12}{\alpha^2} \left\langle \xi_0 \int_0^1 u_0^o d\eta \right\rangle - 6 \int_0^1 F \eta (1 - \eta) d\eta, \quad (4.8)$$

as follows from substitution of (4.6) into (4.5). A similar analysis of the cavity flow provides

$$\langle u_1^c \rangle = -(\alpha H)^2 \left[\frac{\eta}{2} (1 - \eta) \frac{d\langle p_1^c \rangle}{dx} + \int_0^\eta \hat{\eta} \tilde{F} d\hat{\eta} + \eta \left(\int_\eta^1 \tilde{F} d\hat{\eta} - \int_0^1 \eta \tilde{F} d\eta \right) \right], \quad (4.9)$$

$$\langle v_1^c \rangle = -\alpha^2 H^3 \frac{\partial}{\partial x} \left[\frac{\eta^2}{2} \left(\frac{1}{2} - \frac{\eta}{3} \right) \frac{d\langle p_1^c \rangle}{dx} - \frac{1}{2} \int_0^\eta \tilde{F} \hat{\eta}^2 d\hat{\eta} \right. \quad (4.10)$$

$$\left. + \eta \int_0^\eta \tilde{F} \hat{\eta} d\hat{\eta} + \frac{\eta^2}{2} \left(\int_\eta^1 \tilde{F} d\hat{\eta} - \int_0^1 \tilde{F} \eta d\eta \right) \right] + H \int_0^\eta \tilde{G} d\hat{\eta}, \quad (4.11)$$

498 with

$$\frac{d\langle p_1^c \rangle}{dx} = \frac{12}{\alpha^2 H^3} \left\langle \xi_0 \int_0^1 u_0^c d\eta \right\rangle - 6 \int_0^1 \tilde{F} \eta (1 - \eta) d\eta \quad (4.12)$$

499 and

$$\int_0^1 \langle u_1^c \rangle d\eta = -\frac{1}{H} \left\langle \xi_0 \int_0^1 u_0^c d\eta \right\rangle, \quad (4.13)$$

where

$$\tilde{G} = \left(\frac{1 - \eta}{H} \right) \left\langle \frac{\partial \xi_0}{\partial x} \frac{\partial u_0^c}{\partial \eta} \right\rangle - \frac{1}{H^2} \left\langle \xi_0 \frac{\partial v_0^c}{\partial \eta} \right\rangle, \quad (4.14)$$

$$\tilde{F} = \left(\frac{1 - \eta}{H} \right) \left\langle \frac{\partial \xi_0}{\partial t} \frac{\partial u_0^c}{\partial \eta} \right\rangle + \left\langle u_0^c \frac{\partial u_0^c}{\partial x} \right\rangle - \frac{1}{H} \left\langle v_0^c \frac{\partial u_0^c}{\partial \eta} \right\rangle - \frac{2}{\alpha^2 H^3} \left\langle \xi_0 \frac{\partial^2 u_0^c}{\partial \eta^2} \right\rangle. \quad (4.15)$$

500 Using (3.16) together with (4.5) and (4.13) finally gives

$$H \int_{-H}^0 \langle u_1^c \rangle dy = \int_0^1 \langle u_1^o \rangle dy - \frac{1}{2} \text{Re}(\chi) = \frac{1}{2} \text{Re} \left(i\chi \int_0^x \chi^* d\hat{x} \right), \quad (4.16)$$

501 which can be used in conjunction with (3.13) and (3.15) to evaluate the flow rates across
502 the channel $\langle Q_1^o \rangle = \int_0^1 \langle u_1^o \rangle dy \simeq \int_0^1 \langle u_1^o \rangle d\eta$ and cavity $\langle Q_1^c \rangle = \int_{-H}^0 \langle u_1^c \rangle dy \simeq H \int_0^1 \langle u_1^c \rangle d\eta$.

503 To show the complicated structure of the resulting flow, selected results corresponding
504 to a configuration with $\alpha = 6$ and $H = 1.5$ are shown in figures 8(a) ($\mathcal{T} = 0.01$) and 8(b)
505 ($\mathcal{T} = 0.001$). Since the continuity equation, given for the channel in (4.1), contains a
506 source term arising from the membrane deformation, it is not possible to use the stream
507 function to define the streamlines. Instead, the streamlines shown in the upper panels
508 were obtained by direct integration of $dx/\langle u_1 \rangle = dy/\langle v_1 \rangle$. As a consequence, unlike
509 the plots in figure 3, computed with the stream function corresponding to the leading-
510 order harmonic flow, the distance between streamlines in figures 8(a) and 8(b) does not
511 represent the magnitude of the local velocity. A measure of the flow magnitude is provided
512 in this case by the volumetric flow rates shown in the lower panels and also by the color

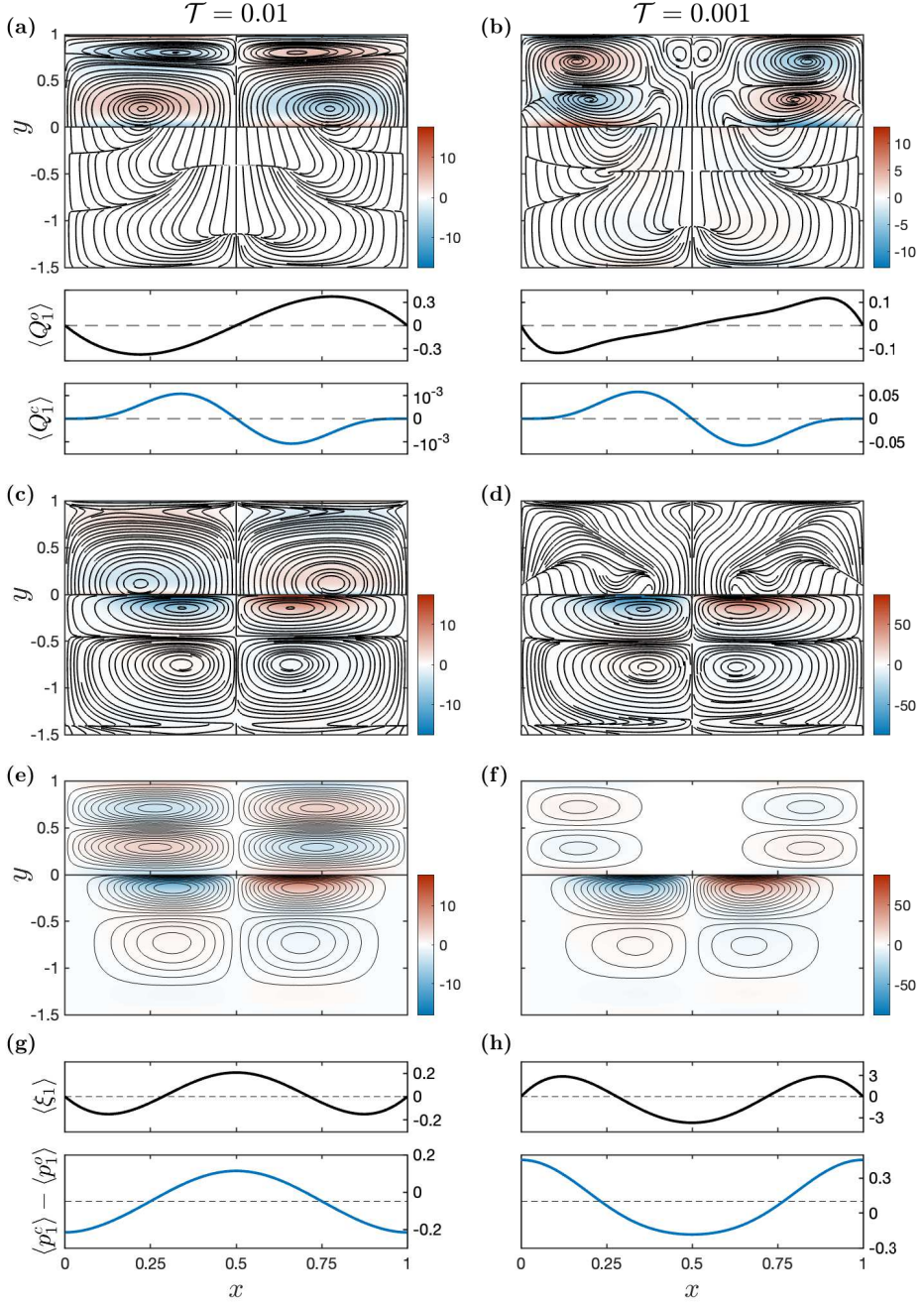


FIGURE 8. Secondary flow for $H = 1.5$ and $\alpha = 6$ with $\mathcal{T} = 0.01$ (left-hand-side plots) and $\mathcal{T} = 0.001$ (right-hand-side plots) including (a,b) streamlines, color contours of vorticity and channel and cavity flow rates corresponding to the steady-streaming velocity $\langle \mathbf{v}_1 \rangle = (\langle u_1 \rangle, \langle v_1 \rangle)$, (c,d) streamlines and color contours of vorticity corresponding to the Stokes-drift velocity $\mathbf{v}_{SD} = (u_{SD}, v_{SD})$, (e,f) streamlines and color contours of vorticity corresponding to the mean Lagrangian velocity $\mathbf{v}_L = \langle \mathbf{v}_1 \rangle + \mathbf{v}_{SD}$, and (g,h) membrane deformation $\langle \xi_1 \rangle$ and stationary transmembrane pressure difference $\langle p_1^c \rangle - \langle p_1^o \rangle$.

513 contours of vorticity $-\partial\langle u_1\rangle/\partial y \simeq -\partial\langle u_1\rangle/\partial\eta$, which are superposed to the streamlines
 514 in the upper panels. As can be seen, for $\mathcal{T} = 0.01$ the motion in the channel is nearly
 515 three orders of magnitude stronger than that in the cavity, while for $\mathcal{T} = 0.001$ their
 516 magnitudes are comparable.

517 The Eulerian flow structure depicted in figures 8(a) and 8(b), symmetric about the
 518 centerline $x = 1/2$, exhibits a variety of singular points. Four centers separated by a
 519 saddle point located along the symmetry plane characterize the flow in the channel for
 520 $\mathcal{T} = 0.01$, with the lower nodes involving streamlines originating at the membrane. As
 521 the membrane tension is increased to $\mathcal{T} = 0.001$, the four centers become spiral points
 522 and the structure becomes more complicated upon the emergence of new saddle points
 523 as well as two new centers. On the other hand, the motion in the cavity is characterized
 524 by the existence of several nodes, giving a flow structure that is markedly different from
 525 that found in the channel.

526 4.2. The mean Lagrangian velocity

527 The complicated streamline structure associated with the steady-streaming velocity
 528 $\langle \mathbf{v}_1 \rangle = (\langle u_1 \rangle, \langle v_1 \rangle)$ shown in figures 8(a) and 8(b) does not represent actual cycle-averaged
 529 trajectories of fluid particles. In characterizing the secondary flow, it is important to
 530 bear in mind that the mean Lagrangian velocity of the fluid particles, smaller than the
 531 oscillatory velocity by a factor ε , has in general a contribution arising from the so-called
 532 Stokes drift (Stokes 1847), additional to that associated with the time-averaged Eulerian
 533 velocity $\langle \mathbf{v} \rangle = \varepsilon \langle \mathbf{v}_1 \rangle$ computed above (see, e.g. Larrieu *et al.* 2009; Alaminos-Quesada
 534 *et al.* 2022, for related channel-flow examples). If the factor ε is incorporated in the scaling
 535 of the Lagrangian velocity $\mathbf{v}_L = (u_L, v_L)$, then it follows that $\mathbf{v}_L = \langle \mathbf{v}_1 \rangle + \mathbf{v}_{SD}$. The
 536 Stokes drift $\mathbf{v}_{SD} = (u_{SD}, v_{SD})$, resulting from small displacements of the Lagrangian
 537 particle during its phase cycle, can be computed from (van den Bremer & Breivik 2018)

$$\mathbf{v}_{SD} = \langle (\delta_x, \delta_\eta) \cdot \nabla \mathbf{v}_0 \rangle, \quad (4.17)$$

538 where $\mathbf{v}_0 = (u_0, v_0)$ is the leading-order oscillatory velocity and (δ_x, δ_η) is the correspond-
 539 ing linear displacement (scaled with ε), to be obtained by integration of the trajectory
 540 equations, with account taken of the coordinate stretching (2.16) in the computation of
 541 the vertical displacement. For example, for the channel the trajectory equations become

$$\frac{\partial \delta_x}{\partial t} = u_0^o \quad \text{and} \quad \frac{\partial \delta_\eta}{\partial t} = v_0^o + (\eta - 1) \frac{\partial \xi_0}{\partial t}, \quad (4.18)$$

yielding upon integration $\delta_x = \int u_0^o dt$ and $\delta_\eta = \int v_0^o dt + (\eta - 1) \xi_0$. Substitution into (4.17)
 provides

$$u_{SD}^o = \frac{\partial}{\partial \eta} \left\langle u_0^o \int v_0^o dt \right\rangle + (\eta - 1) \frac{\partial}{\partial \eta} \langle \xi_0 u_0^o \rangle, \quad (4.19)$$

$$v_{SD}^o = \frac{\partial}{\partial x} \left\langle v_0^o \int u_0^o dt \right\rangle + (\eta - 1) \frac{\partial}{\partial \eta} \langle \xi_0 v_0^o \rangle \quad (4.20)$$

in the channel, while a similar development leads to

$$u_{SD}^c = \frac{\partial}{\partial \eta} \left\langle u_0^c \int v_0^c dt \right\rangle - \left(\frac{\eta - 1}{H} \right) \frac{\partial}{\partial \eta} \langle \xi_0 u_0^c \rangle, \quad (4.21)$$

$$v_{SD}^c = \frac{\partial}{\partial x} \left\langle v_0^c \int u_0^c dt \right\rangle - \left(\frac{\eta - 1}{H} \right) \frac{\partial}{\partial \eta} \langle \xi_0 v_0^c \rangle \quad (4.22)$$

542 in the cavity. It is interesting to note that, just like the steady-streaming velocity $\langle \mathbf{v}_1 \rangle$,
 543 the Stokes velocity is non-solenoidal, as can be seen by computing the divergence to give

$$\frac{\partial u_{SD}^o}{\partial x} + \frac{\partial v_{SD}^o}{\partial \eta} - G = 0 \quad (4.23)$$

544 in the channel, where the function G is defined in (4.2). By way of contrast, the
 545 Lagrangian velocity $\mathbf{v}_L = \langle \mathbf{v}_1 \rangle + \mathbf{v}_{SD}$ is solenoidal, as can be verified by adding (4.23)
 546 to the first equation in (4.1). Correspondingly, the flow rate associated with the Stokes
 547 drift, equal to $\int_0^1 u_{SD}^o d\eta = -\langle \xi_0 \int_0^1 u_0^o d\eta \rangle$ in the channel, balances out with that of the
 548 steady-streaming motion, given for the channel in (4.5), so that the Lagrangian flow rate
 549 satisfies $\int_0^1 u_L d\eta = 0$, as it should.

550 Streamlines computed with use made of (4.19)–(4.22), showing the expected symmetry
 551 about $x = 1/2$, are represented in figures 8(c) and 8(d). According to the above discussion,
 552 corresponding flow rates $\int_0^1 u_{SD}^o dy$ and $\int_{-H}^0 u_{SD}^c dy$ can be obtained by simply changing
 553 the sign of those given for the steady-streaming motion in figures 8(a) and 8(b). Just
 554 as in the case of steady streaming, the resulting flow structure shows multiple singular
 555 points, different in the cavity and in the channel. In contrast, the structure of the mean
 556 Lagrangian flow, depicted in figures 8(e) and 8(f), is somewhat simpler, in that it
 557 comprises four counter-rotating vortices in the channel and in the cavity, resulting in
 558 a zero volume flux, with the flow in the channel displaying symmetry about $y = 1/2$.
 559 As revealed by additional computations, not shown here, the number of Lagrangian
 560 vortices depends on the values of α and \mathcal{T} . For instance, for $\alpha = 6$ and $\mathcal{T} = 10^{-4}$, the
 561 four symmetrically arranged vortices that characterize the channel flow in figures 8(e)
 562 and 8(f) split to give four vortex pairs, each occupying one quadrant of the channel, while
 563 the corresponding cavity flow features in each half space $0 < x < 1/2$ and $1/2 < x < 1$
 564 three dissimilar vortices arranged in a triangular fashion.

565 4.3. Stationary transmembrane pressure difference and membrane deformation

566 While the computation of the oscillatory flow at leading order requires simultaneous
 567 consideration of the membrane deformation, as seen in § 3, the steady-streaming flow
 568 described by (4.6)–(4.12) is independent of the mean membrane displacement $\langle \xi_1 \rangle$. The
 569 computation of $\langle \xi_1 \rangle$ involves the elastic equation (2.12), which yields at this order the
 570 boundary-value problem

$$\mathcal{T} \frac{d^2 \langle \xi_1 \rangle}{dx^2} = \langle p_1^o \rangle - \langle p_1^c \rangle; \quad \langle \xi_1 \rangle(0) = \langle \xi_1 \rangle(1) = 0. \quad (4.24)$$

571 Differentiating once the above equation and substituting (4.8) and (4.12) provides a
 572 third-order equation, which can be integrated with the additional integral condition
 573 $\int_0^1 \langle \xi_1 \rangle dx = 0$, stemming from (2.14), to give

$$\langle \xi_1 \rangle = \frac{1}{\mathcal{T}} \left[x \int_0^1 \mathcal{I}(1-x) dx - 3x(1-x) \int_0^1 \mathcal{I}x(1-x) dx + \int_0^x \mathcal{I}\tilde{x} d\tilde{x} - x \int_0^x \mathcal{I}d\tilde{x} \right] \quad (4.25)$$

574 and

$$\langle p_1^c \rangle - \langle p_1^o \rangle = \mathcal{I}(x) - 6 \int_0^1 \mathcal{I}x(1-x) dx, \quad (4.26)$$

575 where

$$\mathcal{I}(x) = \int_0^x \left[\frac{12}{\alpha^2} \left\langle \xi_0 \int_0^1 (u_0^o + u_0^c/H^3) d\eta \right\rangle + 6 \int_0^1 (F - \tilde{F})\eta(1-\eta) d\eta \right] dx. \quad (4.27)$$

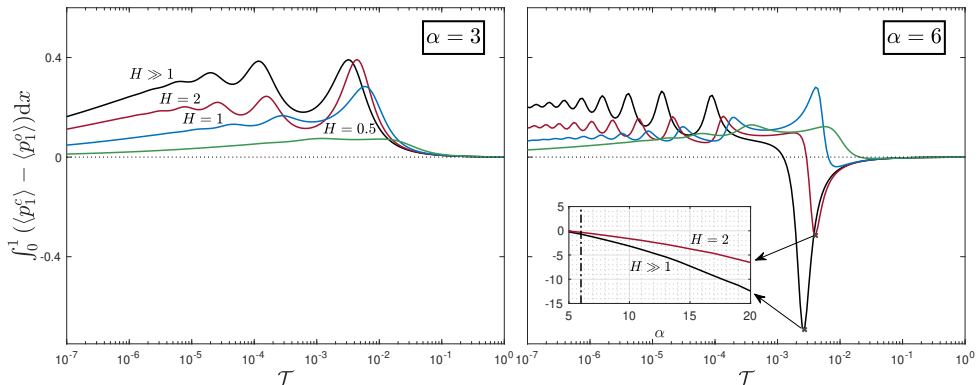


FIGURE 9. The variation with \mathcal{T} of the spatially averaged transmembrane pressure difference $\int_0^1 (\langle p_1^c \rangle - \langle p_1^o \rangle) dx$ for $\alpha = 3$ (left) and $\alpha = 6$ (right) with $H = (0.5, 1, 2, \infty)$. The inset on the right depicts the evolution with α of the peak values of $\int_0^1 (\langle p_1^c \rangle - \langle p_1^o \rangle) dx$ for $H = 2$ (red) and $H = \infty$ (black).

576 The cycle-averaged distributions of membrane displacement $\langle \xi_1 \rangle$ and transmembrane
 577 pressure difference $\langle p_1^c \rangle - \langle p_1^o \rangle$ evaluated from (4.25) and (4.26) with use made of (4.27),
 578 both symmetric about $x = 1/2$, are plotted in figures 8(g) and 8(h). As can be seen, for
 579 $\mathcal{T} = 0.01$, the membrane is convex towards the channel at its center, where the cavity
 580 overpressure reaches its maximum value, while for $\mathcal{T} = 0.001$ the membrane at its center
 581 is concave and the local value of $\langle p_1^c \rangle - \langle p_1^o \rangle$ is negative.

582 A relevant magnitude of interest is the spatially averaged value of the transmembrane
 583 pressure difference

$$\int_0^1 (\langle p_1^c \rangle - \langle p_1^o \rangle) dx = \int_0^1 \mathcal{I}(1 - 6x + 6x^2) dx, \quad (4.28)$$

584 related to the end slope of the membrane $d\langle \xi_1 \rangle/dx(0) = -d\langle \xi_1 \rangle/dx(1)$ according to
 585 $\int_0^1 (\langle p_1^c \rangle - \langle p_1^o \rangle) dx = 2\mathcal{T}d\langle \xi_1 \rangle/dx(0)$, as follows from (4.24). This quantity can be thought
 586 to be representative of the transmural pressure induced by the CSF motion in syringomyelia
 587 cavities, which has been reasoned to play an important role in the develop-
 588 ment of the disease (Bertram 2010; Heil & Bertram 2016; Bertram & Heil 2017), as
 589 SAS overpressures can drive CSF from the SAS through the spinal cord tissue to fill the
 590 cavity. As can be inferred from the pressure distributions in figures 8(g) and 8(h), the
 591 value of $\int_0^1 (\langle p_1^c \rangle - \langle p_1^o \rangle) dx$ is negative for $\mathcal{T} = 0.01$ but positive for $\mathcal{T} = 0.001$, so that
 592 both cavity overpressures and SAS overpressures may arise, depending on the conditions.
 593 Values computed over an extended range of \mathcal{T} for different values of the cavity width
 594 and two different values of α are shown in figure 9.

595 Since the stationary pressure differences originated by the fluid motion are due to
 596 nonlinear interactions involving the leading-order oscillatory solution, the curves in
 597 figure 9 are seen to correlate with those shown in figure 6 for the magnitude of the
 598 oscillating flow rate. Thus, for rigid membranes, corresponding to values of $\mathcal{T} \gtrsim 0.1$, the
 599 stationary pressure differences originated by the fluid motion are negligibly small. The
 600 peak transmembrane pressure difference is attained in figure 9 at an intermediate value
 601 of \mathcal{T} , coincident with the maximum in oscillating flow rate shown in the corresponding
 602 curves of figure 6. Both sets of curves also display oscillations as the membrane develops
 603 a larger number of undulations for $\mathcal{T} = (\lambda_e/L)^4 \ll 1$.

604 As seen in the left-hand-side plot of figure 9, for $\alpha = 3$ the cavity exhibits overpres-

605 sures regardless of the cavity size and membrane tension. However, a more complicated
 606 behavior arises for $\alpha = 6$, a case shown in the right-hand-side plot for which the sign of
 607 $\int_0^1 (\langle p_1^c \rangle - \langle p_1^o \rangle) dx$ depends on the value of H in the intermediate range of values of \mathcal{T}
 608 where the motion is more vigorous. As can be seen, large cavities tend to display negative
 609 pressures, more pronounced for increasing values of H . This aspect of the solution is
 610 further investigated in an inset showing the variation of the peak pressure up to values
 611 of α exceeding the largest value $\alpha = 12$ estimated to be relevant for cardiac-driven CSF
 612 flow in the cervical region.

613 The parametric dependences revealed by figure 9 may have implications regarding the
 614 development of syringomyelia cavities. If one assumes that SAS overpressures are needed
 615 to drive the transmedullary flow responsible for syrinx growth, then, according to the
 616 results shown in the left-hand-side plot, the syrinx would never develop if $\alpha = 3$, since
 617 cavity overpressures (i.e. positive values of $\int_0^1 (\langle p_1^c \rangle - \langle p_1^o \rangle) dx$) prevail for all values of
 618 H and \mathcal{T} . The more complicated variation of $\int_0^1 (\langle p_1^c \rangle - \langle p_1^o \rangle) dx$ shown for $\alpha = 6$ on
 619 the right-hand-side plot suggests that, in the intermediate range of values of \mathcal{T} where
 620 the cavity flow is more pronounced, changes in the syrinx transverse size might have an
 621 important effect, with cavity overpressures turning to SAS overpressures as H increases.
 622 Further increase of H may result in a self-accelerating process that possibly leads to
 623 runaway cavity growth. The curves for $\alpha = 6$ also indicate that, for a constant H ,
 624 there can be situations where the initial SAS overpressure eventually turns to cavity
 625 overpressure as the dimensionless membrane tension \mathcal{T} decreases with increasing syrinx
 626 lengths L , thereby leading to stabilization of a finite-sized syrinx. Naturally, one must
 627 bear in mind that these identified trends pertain to an SAS flow rate of sinusoidal form,
 628 thereby neglecting the influence of higher harmonics, which may have an important effect
 629 on the transmembrane pressure, as discussed below.

630 With the frequency entering in the scale $\rho u_c \omega L$ used to define the dimensionless
 631 pressures p^o and p^c , higher frequencies can be expected to lead to larger transmural
 632 pressures, a trend that is further enhanced by the dependence on $\mathcal{T} \propto \omega^{-2}$ previously
 633 discussed in connection with the curves on figure 6. This observation underscores once
 634 more the potential importance of the higher harmonics arising in the presence of non-
 635 sinusoidal flow rates, as those encountered in the spinal canal. Just as the first or second
 636 harmonic can dominate the sloshing dynamics in the cavity, as revealed by in vivo mea-
 637 surements (Vinje *et al.* 2018) and illustrated in the sample computations of figure 7(b), the
 638 steady transmural pressure difference induced by the higher harmonics can be possibly
 639 larger than that of the fundamental frequency. Because of this frequency-dependent flow
 640 amplification, a result of the underlying FSI dynamics, numerical simulations and in vitro
 641 experiments utilizing a SAS flow rate (or longitudinal pressure gradient) with presumed
 642 sinusoidal waveform may significantly underpredict the associated transmural pressure.

643 To illustrate the previous point, one can use

$$\langle p_1^c \rangle - \langle p_1^o \rangle = \sum_{n=1}^{\infty} n |A_n|^2 (\langle p_{1,n}^c \rangle - \langle p_{1,n}^o \rangle) \quad (4.29)$$

644 to evaluate the streamwise variation of the transmembrane pressure difference $\langle p_1^c \rangle - \langle p_1^o \rangle$
 645 for a channel flow rate of general periodic form (3.24). In the above expression, the
 646 contribution of each mode is weighted by $n |A_n|^2$, where the factor n stems from the
 647 proportionally $\Delta p' \propto \omega$ present in the definition of the dimensionless pressure p . Corre-
 648 spondingly, the dependences on the flow frequency present in the definitions of \mathcal{T} and α
 649 suggest that, in using (4.26) to compute the pressure difference $\langle p_{1,n}^c \rangle - \langle p_{1,n}^o \rangle$ associated
 650 with n th-mode, one must replace \mathcal{T} and α with \mathcal{T}/n^2 and $n^{1/2}\alpha$ when evaluating the

651 integral function (4.27). The expression (4.29) was used to determine the longitudinal
 652 distributions of $\langle p_1^c \rangle - \langle p_1^o \rangle$ shown in figure 7(c), corresponding to the sinusoidal and
 653 complex-wave flow rates shown in figure 7(a). As anticipated in the previous paragraph,
 654 the presence of higher harmonics in the channel-flow wave form has a dramatic effect
 655 on the magnitude of $\langle p_1^c \rangle - \langle p_1^o \rangle$. Correspondingly, the spatially averaged transmembrane
 656 pressure difference, which takes the values $\int_0^1 (\langle p_1^c \rangle - \langle p_1^o \rangle) dx = (-0.0117, -0.0151)$ for
 657 $H = (1, 4)$ when a sinusoidal SAS-flow rate is assumed, increases to $\int_0^1 (\langle p_1^c \rangle - \langle p_1^o \rangle) dx =$
 658 $(0.1404, -0.3059)$ for $H = (1, 4)$ when the physiologically correct flow rate is used in
 659 the computation. For the latter, the change in sign of the transmembrane pressure with
 660 increasing H may have implications concerning transmedullary flow. Clearly, additional
 661 work involving more accurate models is needed before these identified trends can be used
 662 for predictive purposes.

663 5. Conclusions

664 The time-periodic hydrodynamics of syringomyelia cavities, involving a FSI problem
 665 in which the motion in the spinal-cord cavity is coupled with that in the surrounding
 666 subarachnoid space through the dynamic response of the separating tissue, has been
 667 analyzed with use of a canonical flow configuration, schematically represented in fig-
 668 ure 2(c). In seeking maximum simplification, the conservation equations are written in
 669 the slender-flow approximation, appropriate for the description of long syrinxes, with
 670 the separating tissue represented by a membrane satisfying a linearly elastic equation.
 671 An asymptotic analysis for small stroke lengths leads to closed-form expressions for the
 672 velocity, pressure and membrane displacement, involving integrals that can be easily
 673 evaluated to investigate the characteristics of the solution for relevant values of the three
 674 controlling parameters, namely, the Womersley number α , the reduced membrane tension
 675 \mathcal{T} , and the cavity-to-channel width ratio H .

676 The oscillatory flow that appears at leading order, with zero mean, characterizes
 677 the sloshing motion in the cavity. An important finding of the analysis is that, as
 678 a consequence of the underlying FSI dynamics, the magnitude of the cyclic motion
 679 induced in the cavity by the external flow oscillations exhibits a strong dependence
 680 on the driving frequency. Because of this frequency-dependent flow amplification, in
 681 systems involving non-sinusoidal external flow, the intracavitary flow may be dominated
 682 by higher harmonics. For example, for the flow-rate waveform encountered in the spinal
 683 canal, shown in figure 7(a), it was found that the flow in the cavity may exhibit
 684 multiple pulsations per cycle (see figure 7(b)), in agreement with previous *in vivo*
 685 observations pertaining to flow in syringomyelia cavities (Vinje et al. 2018). Interestingly,
 686 also consistent with those observations, the model predicts that the number of pulsations
 687 per cycle decreases as the transverse dimension of the cavity shrinks. It is worth noting
 688 that the current prediction is based on a linear elastic model, and therefore precludes
 689 effects of nonlinear cavity resonance, which should be investigated in future work.

690 The first-order corrections are seen to include a steady component resulting from
 691 the combined action of the convective acceleration and the nonlinear interactions of
 692 the membrane deformation with transverse velocity gradients. The sum of the steady
 693 streaming and the Stokes drift determines the recirculating mean Lagrangian motion, as
 694 depicted in figure 8. The associated cycle-averaged transmembrane pressure difference
 695 $\langle p_1^c \rangle - \langle p_1^o \rangle$, which represents in the model the stationary transmural pressure driving the
 696 CSF transmedullary flow in syringomyelia cavities, has been computed over extended
 697 parametric ranges. The results reveal that, just like the leading-order oscillatory flow,

698 the transmembrane pressure difference shows a prominent dependence on the frequency,
 699 once more underlying the potential relevance of higher harmonics. Depending on the
 700 conditions, the cycle-averaged intracavitary pressure can be either higher or lower than
 701 the SAS pressure. For the sinusoidally varying flow rate of figure 9, large SAS overpres-
 702 sures (negative values of $\langle p_1^c \rangle - \langle p_1^q \rangle$) are predicted when $\alpha \gtrsim 6$ for large cavities in the
 703 intermediate range of values of \mathcal{T} for which the sloshing motion is more pronounced.
 704 These large SAS overpressures and their potential contribution to the transmedullary
 705 flow clearly warrant future investigation.

706 Future extensions of the analytical work presented here should consider an improved
 707 model for the dynamics of the tissue separating the cavity from the SAS, possibly
 708 replacing the elastic membrane with a compliant wall having inertia, damping, and
 709 flexural rigidity (Davies & Carpenter 1997). Axisymmetric configurations (i.e. a fluid-
 710 filled tubular cavity separated from a coaxial channel by a flexible membrane) are
 711 attractive for investigations of canalicular syringomyelia. In this axisymmetric geometry,
 712 the restoring force arises primarily from the hoop stresses induced by the azimuthal
 713 stretching, so that (2.12) would be replaced with the condition that the membrane
 714 displacement be linearly proportional to the transmembrane overpressure, with axial
 715 membrane tension becoming important only inside boundary-layer regions located at the
 716 two ends of the cavity. While the quantitative results of the axisymmetric model can be
 717 expected to depart from those of the 2D cavity, the solution would probably exhibit many
 718 of the features identified above, including the strong dependence of the cavity flow on
 719 the frequency of the external oscillatory stream and the existence of a steady transmural
 720 pressure.

721 More accurate models accounting for the finite thickness of the separating tissue and
 722 its poroelastic properties (Venton *et al.* 2017; Cardillo & Camporeale 2021) would be
 723 needed to enable accurate quantitative predictions. A thorough investigation of effects
 724 of flow-rate waveform could help further assess effects of higher harmonics. Also, by
 725 modifying the width distribution along the channel representing the SAS, the model could
 726 be readily extended to address effects of SAS tapering and stenosis, which are known to
 727 lead to important changes in the flow (Bertram 2010; Martin *et al.* 2010; Heil & Bertram
 728 2016; Bertram & Heil 2017). The results of the theoretical analysis can help guide future
 729 computational efforts aimed at providing accurate quantitative predictions of transmural
 730 pressure differences, required to clarify outstanding questions pertaining to the “filling
 731 mechanism” (Stoodley 2014). In view of the present results, besides consideration of
 732 anatomically correct models, these future computations should consider CSF flow-rate
 733 waveforms and spinal-cord elastic properties that are physiologically correct, as needed
 734 for an accurate description of high-frequency transmural flow amplification.

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