

On the role of aortic valve architecture for physiological haemodynamics and valve replacement. Part II: spectral analysis and anisotropy

Pascal Corso (✉ pascal.corso@unibe.ch)

University of Bern <https://orcid.org/0000-0001-7875-1080>

Dominik Obrist

University of Bern

Research Article

Keywords: Aortic stenosis, Bioprosthetic aortic valve, Flow turbulence, Fluid-Structure Interaction simulations, Kinetic energy anisotropy, Spectral analysis, Valve design

Posted Date: September 27th, 2023

DOI: <https://doi.org/10.21203/rs.3.rs-3387570/v1>

License: © ⓘ This work is licensed under a Creative Commons Attribution 4.0 International License.

[Read Full License](#)

1 **On the role of aortic valve architecture for physiological**
2 **haemodynamics and valve replacement. Part II: spectral analysis**
3 **and anisotropy.**

4 Pascal Corso¹, Dominik Obrist¹

5 **Affiliations:** ¹ARTORG Center for Biomedical Engineering Research, University of Bern,
6 Bern, Switzerland.

7 **Correspondence:** Pascal Corso, University of Bern, Bern, Switzerland.

8 E-mail: pascal.corso@unibe.ch

9 Abstract

10 Severe aortic valve stenosis can lead to heart failure and aortic valve replacement (AVR) is the
11 primary treatment. However, increasing prevalence of aortic stenosis cases reveal limitations
12 in current replacement options, necessitating improved prosthetic aortic valves. In this study,
13 we investigate flow disturbances downstream of severe aortic stenosis and two bioprosthetic
14 aortic valve (BioAV) designs using advanced energy-based analyses. Spectral analysis shows
15 kinetic energy (KE) decay variations, with the stenotic case aligning with Kolmogorov's
16 theory, while BioAVs differ. We explore the impact of flow helicity on KE transfer and
17 decay in BioAVs. Probability distributions of modal KE anisotropy unveil flow asymmetries
18 in the stenotic and one BioAV case. Moreover, an inverse correlation between modal KE
19 anisotropy and normalised helicity intensity is noted, with the coefficient of determination
20 varying among the valve configurations. Leaflet dynamics analysis highlights a stronger
21 correlation between flow and biomechanical KE anisotropy in one BioAV due to higher leaflet
22 displacement magnitude. These findings emphasise the role of valve architecture in aortic
23 turbulence and its significance for BioAV performance and energy-based design optimisation.

24 Main

25 Calcific aortic valve stenosis is characterised by a progressive deterioration, remodelling
26 and thickening of the native aortic valve leaflet tissue, causing a reduction in its functional
27 flexibility. This condition, referred to as aortic stenosis (AS), results in increased resistance
28 to blood flow from the left ventricle to the aorta, particularly during systole and the potential
29 for blood regurgitation during diastole^{6,7,29}. The long-term implications of this pathology
30 are serious. The AS is indeed generally defined as severe when a significant left ventricular
31 outflow tract (LVOT) obstruction leading to a reduced orifice area and high downstream
32 jet velocity is present and symptoms such as dyspnea, heart failure, chest pain or syncope
33 appear³. To address this critical issue, the replacement with valvular prostheses has emerged
34 as a prevalent solution. These prostheses come in two primary types: mechanical heart
35 valves made from rigid materials such as titanium or carbon and bioprosthetic or tissue
36 aortic valves (BioAVs) manufactured based on biological tissue. The replacement of the
37 diseased native aortic valve is achieved through a medical procedure known as aortic valve
38 replacement (AVR). Aortic valve stenotic disease is the most commonly occurring valvular
39 pathology in developed countries (afflicting 9 million people worldwide) and its prevalence
40 has been increasing with population ageing¹. Moreover, surgical aortic valve replacements
41 tally around 300,000 cases annually and this number is projected to double by 2050 due to
42 the ageing global population²¹.

43 Previous studies have extensively explored the haemodynamic performance of aortic
44 valves made from biological materials like porcine or bovine pericardium^{2,4,6,13,27,29}. How-
45 ever, the correlation between the kinetic energy (KE) present within the valve components
46 (i.e. leaflets and supporting ring) and the energy carried by the turbulent structures in the
47 flow surrounding the BioAV has yet to be investigated, either *in vitro* or *in silico*.

48 In their work, Bescek *et al.*⁴ presented a broad computational characterisation of the tur-
49 bulent features of the flow downstream of one bioprosthetic aortic valve model under peak
50 systolic conditions. One-dimensional wavenumber energy spectra were calculated at various
51 distances from the sino-tubular junction and it was argued that the spectra based on points

52 diametrically aligned where the turbulent dissipation rate is the largest could be connected
53 to shear-induced thrombocyte activation. Finally, it was noticed the presence of zones with
54 elevated and fluctuating wall shear stress at the aortic wall that could possibly underline the
55 presence of endothelial lesions in these zones. Nonetheless, the wavenumber kinetic energy
56 spectra considered in Becsek *et al.*⁴ were one-dimensional and limited to a confined region of
57 the flow along a line perpendicular to the centerline, thus excluding the study of anisotropy
58 in the kinetic energy (KE).

59 Besides, in Corso *et al.*^{6,7,8}, similar conclusions were drawn using both *in vitro* three-
60 dimensional particle tracking velocimetry experiments and from direct numerical simulation
61 downstream of a stenotic aortic valve as to the detrimental effect of turbulence on blood
62 platelet damage and on the production of important pressure loss due to elevated haemody-
63 namic turbulent stresses. Moreover, it was emphasised that taking into account Reynolds'
64 stress close to the wall for the accurate evaluation of the wall shear stress from flow field
65 data with limited spatial resolution is crucial when disturbed aortic flows are considered.

66 Recently, Gallo *et al.* have explored the relationship between phase-averaged and fluctu-
67 ating helicity and phase-averaged and turbulent kinetic energy for one model of mechanical
68 heart valve prostheses and for one model of bioprosthetic valve. To this end, they simu-
69 lated the coupled fluid-structure interaction problem by applying the interface conditions
70 through an immersed body surface method. It was found in the study that haemodynamics
71 downstream of the mechanical valve presenting larger phase-averaged and fluctuating helicity
72 than that downstream of bioprosthetic valve. For both heart valve types, strong linear cor-
73 relations were found between volume-averaged kinetic energy and helicity when considering
74 phase-averaged or fluctuating quantities. Peaks of turbulent kinetic energy or fluctuating he-
75 licity for both heart valve types was delayed as compared to the peaks of mean kinetic energy
76 and phase-averaged helicity. Despite this study introduces novelty regarding the relationship
77 between flow helicity and kinetic energy and despite the use of 20 simulated cardiac cycles
78 to compute flow field statistics, several limitations should be recognized. These include the
79 use of a simplified mass-spring model to solve the dynamics of the deformable leaflets, the

80 assumption of a rigid aortic wall and the absence of a ring supporting the valve leaflets in
81 the simulations. Furthermore, an in-depth examination of the spatial distribution of kinetic
82 energy, helicity, and their relationship to leaflet and valve designs was not established.

83 The present work is the second part of a comprehensive two-part study. Both parts utilise
84 validated and high-fidelity computational approaches. In this paper, the analysis of blood
85 motion strives to comprehensively characterise the turbulence by inspecting the energy car-
86 ried by the vortical structures of the flow following vortex ring formation, a shedding process
87 including vortex stretching and advection⁹. We seek to observe the typical energy decay of
88 kinetic energy (KE) as described in the theory of turbulence²³. Additionally, we investigate
89 any deviations from this theoretical energy decay along with examining the dissipation rate
90 of KE. Connection between leaflet geometries and turbulence characteristics is underlined. In
91 addition, we analyse and correlate the spatial distributions of the newly introduced concepts
92 of kinetic energy anisotropy and helicity intensity over spherical shells close to the valvular
93 orifice. There has been no previous computational study and comprehensive energy-based
94 analyses that include valve bioprostheses of different designs alongside a comparative assess-
95 ment with a severe stenotic case. This work significantly contributes to the development
96 of optimally designed valves by thoroughly elucidating the flow-energy-based mechanisms
97 downstream of BioAVs in comparison to those encountered downstream of a severe aortic
98 stenosis.

99 **Spectral analysis and turbulence characteristics**

100 The log-log graphs presented in Fig. 1 (a, b, c) display the distribution of the modal KE
101 at different scalar angular wavenumbers for various time instances considered over systole,
102 represented by thin grey lines. Details on the geometry of three valvular configurations, the
103 stenosed aorta case and the two newly designed bioprosthetic valves (VLth30 and Ulth0),
104 are presented in the Methods section of the first part of the study⁹.

105 In the stenotic case, we note that the trend in the decay of energy as a function of the

106 wavenumber varies depending on the time instant considered as shown by the large dif-
 107 ference in E_{u^2} between the dash-dot black line and the solid thick black line, especially for
 108 wavenumbers larger than 2,000 rad/m . The maximum non-dimensional modal KE curve (i.e.
 109 solid thick line in Fig. 1 (a)) corresponds to the wavenumber spectra at instants $t=0.121$ s,
 110 0.142 s and 0.172 s. The minimum non-dimensional modal KE curve (cf. dash-dot line in
 111 Fig. 1 (a)) corresponds to the spectra at $t=0.1$ s.

112 Concerning the VLth30 BioAV case, the log-log plot (see Fig. 1 (b)) shows that the variations
 113 in the energy decay over the different time instants is smaller than that noted in the stenotic
 114 case. These differences between time instances in E_{u^2} are even smaller in the Ulth0 BioAV
 115 case. Indeed, as presented in the first part of the current comprehensive study⁹, the Ulth0
 116 valve design leads to very limited leaflet motion after their opening throughout systole and
 117 the jet shape downstream remains relatively stable, except for Kelvin-Helmholtz instability
 118 (KHI) arising between $t=0.12$ s and $t=0.21$ s.

119 In order to verify whether the logarithmic slope of $-5/3$ characteristic of the inertial subrange
 120 for turbulent flows according to Kolmogorov's theory²³ can be noted, E_{u^2} has been divided
 121 by $\kappa^{-5/3}\varepsilon^{2/3}$ and the graphs of the first-order derivative with respect to the wavenumber
 122 are presented in Fig. 1 (d, e, g). Additionally, the temporal evolution of the turbulent KE
 123 dissipation ε is presented in Fig. 2 (b).

124 For the stenosed aorta case, the curves of minimum, median and maximum non-dimensional
 125 $d\left(\frac{E_{u^2}}{\kappa^{-5/3}\varepsilon^{2/3}}\right)/d\kappa$ are almost superimposed on each other. Moreover, these curves follow
 126 the universal scaling law predicted by Kolmogorov turbulence theory over a wide range of
 127 wavenumbers, i.e. between 2,000 and 20,000 rad/m where $d\left(\frac{E_{u^2}}{\kappa^{-5/3}\varepsilon^{2/3}}\right)/d\kappa$ is equal to 0.
 128 The self-similarity of vortical structures in the inertial subrange is then, in the stenotic
 129 case, observed throughout the whole systolic time interval under consideration. It is worth
 130 mentioning that, for the computation of the wavenumber energy spectra, the mode of the
 131 log-normal distribution of the correlation function over the points on spherical shells in
 132 Fourier space has been taken as the most representative energy level at the considered scalar
 133 wavenumber.

134 In the case of the VLth30 bioprosthesis, the power-law with the exponent of -5/3 character-
 135 istic of the inertial subrange for the decay of energy does not appear, except at time instants
 136 $t = 0.1$ s, 0.11 s, 0.12 s and 0.245 s, over a narrow range of wavenumbers (see the video in
 137 the supporting information). Conversely, peaks of energy at $\kappa = 5,000$ *rad/m* and $\kappa = 7,500$
 138 *rad/m* are noticeable as exhibited in Fig. 1 (e). For values of κ ranging from $1,000$ to $6,500$
 139 *rad/m*, it has been calculated that the minimum, median and maximum values in time of
 140 the $E_u^2/\varepsilon^{2/3}$ curve scale as $\kappa^{-5/3} \ln(\kappa)$, $\kappa^{-5/3} \ln(\kappa)$ and $\kappa^{-7/6}$, respectively.
 141 In the Ulth0 BioAV case, the curve of $d\left(\frac{E_u^2}{\kappa^{-5/3}\varepsilon^{2/3}}\right)/d\kappa$ equals 0 over wavenumbers ranging
 142 from $4,000$ to $9,000$ *rad/m* for time instances between $t = 0.12$ s and $t = 0.19$ s. Interestingly,
 143 these instants correspond to the times at which the KHI establishes as discussed in⁹. In Fig. 1
 144 (f), we observe that, for scalar wavenumbers from $1,000$ and $6,000$ *rad/m*, the minimum,
 145 median and maximum in time of $\frac{E_u^2}{\varepsilon^{2/3}}$ curve scale as κ^{-2} , $\kappa^{-4/3}$ and $\kappa^{-3/2}$, respectively.

146 [Figure 1 about here.]

147 Fig. 2 (a) presents the time evolution of the Kolmogorov length scale derived from the
 148 dissipation rate of turbulent KE ε computed over the spherical shells (cf. Fig. 8). This length
 149 scale is the smallest in the stenotic case with a value of about 20 μm constant over systole.
 150 In the cases of the two valvular bioprostheses, the Kolmogorov length scale is two to four
 151 times the value evaluated downstream of the stenosed aorta. Furthermore, as a consequence
 152 of the peak in energy dissipation at $t=0.15$ s (Fig. 2 (b)), the Ulth0 case exhibits a reduction
 153 in the Kolmogorov length scale, decreasing from 80 to 40 μm . The Kolmogorov length scale
 154 in the VLth30 case ranges from 40 to 50 μm over systole.

155 The integral length scale \mathfrak{L} is presented in Fig. 2 (c) and was determined by computing
 156 the autocorrelation function for each component of the velocity vector \mathbf{u} in physical space.
 157 Subsequently, an equivalent autocorrelation function was calculated over points distributed
 158 on spherical surfaces and \mathfrak{L} was obtained by computing the L^2 -norm of this equivalent
 159 autocorrelation function in each direction. In the stenotic case, the time-averaged integral
 160 length scale is approximately 2.8 *mm* while for the two BioAV prostheses, the time-averaged
 161 \mathfrak{L} ranges from 4.5 to 5 *mm*.

[Figure 2 about here.]

162

163 Energy distribution

164 In this section, we investigate the distribution of energy intensity in physical space, i.e. over
165 spherical surfaces, as defined in the Materials and Methods section. We intend to compare
166 the energy levels among the following cases: the stenosed aorta case, the two newly designed
167 bioprosthetic valves (VLth30 and Ulth0) and a bioprosthetic case presented in the study by
168 Gallo *et al.*¹³.

169 With regard to the turbulent KE intensity, we note in Fig. 3 (a) and (e) that the maximum
170 $\mathbb{I}_{u'^2}$ for the three valvular bioprostheses represents 30% of the maximum $\mathbb{I}_{u'^2}$ in the stenosis
171 case. The time-averaged fluctuating energy intensity for the three BioAV cases is one-sixth
172 that of the stenotic case as illustrated in Fig. 4 (e). We also observe from Fig. 4 (a) that
173 the temporal profile shape for the Ulth0 BioAV case is congruent with the BioAV case
174 investigated in Gallo *et al.*, with the exception that, due to differences in the acceleration
175 and deceleration inflow conditions, the green curve of the BioAV studied by Gallo *et al.* is
176 shifted in time relative to the Ulth0 curve.

177 In regards to the fluctuating enstrophy intensity $\mathbb{I}_{\omega'^2}$, the time-averaged value for the VLth30
178 valvular case is thrice that of the stenotic case and the Ulth0 BioAV case. The elevated levels
179 of fluctuating enstrophy in the VLth30 case arise from the non-axisymmetric and more
180 pronounced leaflet motion, with a displacement magnitude of approximately 2 mm during
181 systole^{27,9}. This finding is consistent with the higher levels of streamwise instantaneous
182 vorticity highlighted in the first part of the study⁹. For the VLth30 case, it is worthwhile
183 to note that the times at which a local minimum in the fluctuating enstrophy temporal
184 evolution is found (i.e. t=0.12, 0.156, 0.21, 0.256 s) correspond to the times at which the
185 area at the vena contracta is maximum⁹.

186 Closely related to the notion of vortex stretching is helicity in flows. Helicity is the integral
187 over a volume of interest of the inner product between velocity and vorticity vectors. Helicity

188 plays an important role in the generation and evolution of vortices and it was shown that it
189 tightly connects to the knottedness and the twisting of vortex lines^{18,19}.

190 In addition, helicity is known to inhibit the transfer of energy towards smaller scales, since
191 the statistical alignment of velocity and vorticity leads to partial suppression of the non-linear
192 term and to a relatively low dissipation¹³. The results in Fig. 3 (c, d, e) are aligned with this
193 statement. For both the VLth30 and Ulth0 cases, the time-averaged and maximum values
194 of unsigned helicity intensity (III and IV) are 2 to 2.5 times as large as those observed in the
195 stenotic case. Moreover, the curves of signed helicity intensity (Fig. 3(d)) for the VLth30
196 and Ulth0 BioAVs exhibit peaks with a magnitude 10 times larger than those observed in
197 the stenotic case and in the BioAV case investigated in Gallo *et al.*. As emphasised in
198 the previous section, the logarithmic slope of the three-dimensional wavenumber spectra
199 (represented by the median, maximum, and minimum curves over time) for the two BioAV
200 cases is larger than the -5/3 logarithmic slope predicted by Kolmogorov for isotropic and
201 homogeneous turbulence, which is found to hold though in the stenotic case. This suggests
202 an inhibition of non-linear advection in the energy cascade, likely due to the presence of
203 local helical flow motion, in the region near the bioprosthetic valvular orifice. Finally, it is
204 worth noting in Fig. 3(d) that the signed helicity intensity $\mathbb{I}_{h'2}$ in the VLth30 case exhibits
205 an average negative value across spherical shells between $t=0.1$ and $t=0.15$ s. This indicates
206 the prevalence of counter-clockwise helical structures in the vicinity of the valvular orifice
207 during this time period. In contrast, in the Ulth0 case, the signed helicity intensity remains
208 positive until $t=0.2$ s, suggesting the dominance of clockwise helical motion. After $t=0.15$ s,
209 a decrease in $\mathbb{I}_{h'2}$ is observed, eventually reaching small negative values until $t=0.28$ s.

210 [Figure 3 about here.]

211 Anisotropy in the modal kinetic energy and helicity in- 212 tensity

213 In this section, the novel quantities defined in Eq. 27 and Eq. 28 representing the modal
214 KE anisotropy $\mathbb{I}_{p_s}^{\text{flow anis}}$ and normalised helicity intensity $\mathbb{I}_{p_s}^{h|\text{sph}}$, respectively, are characterised
215 and correlated through probability density function (PDF) and spatial heatmaps, unrolled
216 by means of a cylindrical projection from the distribution over spherical shells described in
217 the Materials and Methods section.

218 In Fig. 4 (a), the log-normal PDF fitted to the absolute value of $\mathbb{I}_{p_s}^{\text{flow anis}}$ distributed over the
219 spherical surfaces (cf.Fig.8) and evolving in time is presented. We observe that, across all
220 time instances considered between $t=0.1$ and 0.3 s, the mode of the log-normal distribution
221 for $|\mathbb{I}_{p_s}^{\text{flow anis}}|$ in the stenotic case is 35% and 65% higher than the modes of the log-normal
222 distributions in the VLth30 and Ulth0 cases, respectively. In the same figure, the modes for
223 each time instance considered are represented with lozenges and the stenotic case displays
224 a maximum mode value of 9.5% against 7.9% for the VLth30 case and 6% for the Ulth0
225 case. We can also notice this trend in the PDFs and boxplots of $\mathbb{I}_{p_s}^{\text{flow anis}}$ shown in Fig. 4 (b)
226 for the three valvular cases. In fact, the maximum values of $\mathbb{I}_{p_s}^{\text{flow anis}}$ equals 136%, 92% and
227 81.3% for the stenotic, VLth30 and Ulth0 cases, respectively. In Fig. 4 (b), it can be seen
228 that the median value for $\mathbb{I}_{p_s}^{\text{flow anis}}$ is positive for the two bioprosthesis cases, with a value
229 of around 1.5% whereas the median value for the stenotic is negative and equal to -2.2%.
230 The asymmetry in the probability distribution of modal KE is indicative of asymmetries
231 in jet flows, which are associated with complex vortex dynamics⁹, especially when helicity
232 is non-zero¹⁰. The skewness of the VLth30 probability density function (PDF) is negative,
233 while that of the stenotic PDF is positive. Both cases exhibit a similar interquartile range
234 (IQR) value of 35%. In contrast, the PDF for the Ulth0 case appears almost symmetric,
235 as emphasised by Corso et al.⁹, as a consequence of a triangular jet flow topology with low
236 eccentricity distance throughout systole. The IQR in the Ulth0 case equals 27%.

237 [Figure 4 about here.]

238 In Fig. 5, the spatial distribution of the absolute value of the modal KE anisotropy
 239 intensity $|\mathbb{I}_{p_s}^{\text{flow anis}}|$ (Eq. 27) is presented by unrolling the spherical through a cylindrical
 240 projection. The standard deviation σ over the investigated time instances and the time
 241 average ($\bar{\cdot}$) of $|\mathbb{I}_{p_s}^{\text{flow anis}}|$ are compared based on the two-dimensional (α, φ) maps.
 242 In the stenotic case, in Fig. 5 (c), we observe that $|\overline{\mathbb{I}_{p_s}^{\text{flow anis}}}|$ reaches its maximum, with
 243 values ranging from 25% to 40%, at the azimuthal angles corresponding to the locations
 244 of commissures 1, 2, and 3 (i.e., at $\alpha \approx -120^\circ, -10^\circ$ and 80°). The standard deviation
 245 $\sigma(|\mathbb{I}_{p_s}^{\text{flow anis}}|)$ attains peak values of 15% in the region between commissures 1 and 3. This
 246 region corresponds to a zone where low flow velocities and few coherent vortical structures
 247 with limited stretching are found⁹. We also notice a region with moderately high values of
 248 5% for $\sigma(|\mathbb{I}_{p_s}^{\text{flow anis}}|)$ at elevation angles $\varphi > 50^\circ$.
 249 Concerning the VLth30 BioAV, Fig. 5 (e) and (f) reveal that $\sigma(|\mathbb{I}_{p_s}^{\text{flow anis}}|)$ and $|\overline{\mathbb{I}_{p_s}^{\text{flow anis}}}|$ are
 250 the largest for $\varphi < -50^\circ$. In addition, peak values in the standard deviation of absolute value
 251 of modal KE anisotropy, reaching 15%, are found at azimuthal angles of $-150^\circ, -30^\circ$ and 90° .
 252 We observe that these peaks are aligned with the position of the three posts of the BioAV
 253 ring (see Fig.2 of the paper discussing the first part of the study for the description of the
 254 BioAV geometries⁹).
 255 With respect to the Ulth0 bioprosthesis, the values for the spatial distribution of $\sigma(|\mathbb{I}_{p_s}^{\text{flow anis}}|)$
 256 and $|\overline{\mathbb{I}_{p_s}^{\text{flow anis}}}|$ are noticeably lower, as previously indicated based on the PDFs and the modes
 257 of the fitted log-normal distribution. However, peaks in the time-averaged absolute value of
 258 modal KE anisotropy $|\overline{\mathbb{I}_{p_s}^{\text{flow anis}}}|$ can be observed in Fig. 5 (i). Akin to the VLth30 case, these
 259 peaks are positioned at angles α aligned with the positions of the valvular ring posts.

260 [Figure 5 about here.]

261 Fig. 6 presents the comparison and point-to-point correlation between the unrolled maps
 262 of time-averaged modal KE anisotropy and normalised helicity intensity. The interest in
 263 conducting such a correlation lies in the findings presented by Gallo *et al.*¹³, which reveal
 264 a strong linear relationship between phase-averaged and fluctuating helicity and KE. In the

265 study by Gallo *et al.*, the correlation was established based on volume-averaged energy and
 266 helicity over the cardiac cycle.

267 In Fig. 6 (b, f, j), the heatmaps of time-averaged normalised helicity intensity are displayed.

268 These maps suggest that the spatial distribution of $\overline{\mathbb{I}_{p_s}^{h|\text{sph}}}$ is inversely proportional to the
 269 spatial distribution of $|\overline{\mathbb{I}_{p_s}^{\text{flow anis}}}|$, as depicted in the maps shown in Fig. 6 (a, e, i). In order

270 to conduct a point-to-point correlation for the three valvular cases, a non-linear least-square
 271 regression problem is solved. To achieve this, the points on the (α, φ) maps are divided into

272 two sets: one is the training set used for fitting the coefficients A and B in the power-law
 273 equation of the form $|\overline{\mathbb{I}_{p_s}^{\text{flow anis}}}| = A \left[\overline{\mathbb{I}_{p_s}^{h|\text{sph}}} \right]^B$ and the other is the testing set used to evaluate

274 the prediction accuracy. The latter is assessed through the coefficient of determination
 275 (R^2)^{7,8,27}. From Fig. 6 (c, g, k), it can be noted that, for the three valve configurations,

276 $|\overline{\mathbb{I}_{p_s}^{\text{flow anis}}}|$ inversely correlates with $\overline{\mathbb{I}_{p_s}^{h|\text{sph}}}$ as demonstrated by the negative fitted exponent B
 277 (cf. Table 1 in the supplementary information). The accuracy of the regression from R^2

278 evaluated on the training (Fig. 6 (g)) and testing (Fig. 6 (h)) data points of the heatmaps
 279 is the highest with a value of about 0.75 in the VLth30 BioAV case. This valve design leads

280 to non-axisymmetric leaflet motions with displacement magnitude of about 2 mm during
 281 systole (see supplementary material of⁹). The eccentricity of the jet calculated in a proximal

282 cross-section as well as the area at the vena contracta is also evaluated in⁹ and it has been
 283 shown that the leaflet motions produce stronger levels of vortex stretching magnitude as

284 compared to the Ulth0 BioAV case, whose leaflets are almost immobile throughout systole.
 285 The latter case exhibits a low coefficient of determination R^2 of 0.1 based on both the

286 training and testing data points from the heatmaps or spherical shells, indicating a weaker
 287 anti-correlation between modal KE anisotropy and normalised helicity intensity. Regarding

288 the stenotic case, the correlation accuracy on the training dataset is moderate with $R^2 =$
 289 0.43 (see Fig. 6 (c)) while the coefficient of determination drops to 0.1 on the testing data

290 points (Fig. 6 (d)).

291 The variation in the prediction accuracy and strength of correlation among the three valvular
 292 cases under examination is an interesting point to analyse. In fact, as stated by Gallo *et*

293 *al.*¹³, the role of helicity in the generation and evolution of turbulence may depend on the
 294 topology of the flow and vortices, namely dependent on the vortex stretching process. In the
 295 first part of this broad study⁹, we highlight three different jet flow topologies connected to
 296 the valvular orifice architecture. It can then be postulated that flow asymmetries introduced
 297 by moving leaflets, as highlighted in the VLth30 case by the presence of a jet with high
 298 velocities in a proximal cross-section⁹, which changes shape over time and exhibits varying
 299 eccentricity, result in higher levels of KE anisotropy close to the orifice and a stronger
 300 negative correlation with normalised helicity intensity. Finally, it is worth noting for both
 301 BioAV cases, similarly to what was observed for $\overline{|\mathbb{I}_{p_s}^{\text{flow anis}}|}$ and $\sigma(|\mathbb{I}_{p_s}^{\text{flow anis}}|)$, high values of
 302 $\overline{\mathbb{I}_{p_s}^{\text{lsph}}}$ are found at azimuthal angles where the three valve ring posts (indicated by the letter
 303 P in Fig. 6) are located.

304 [Figure 6 about here.]

305 **Leaflet dynamics analysis and anisotropy in the valve**

306 **motion**

307 Since we previously highlighted the potential impact of moving leaflets on the degree of
 308 correlation between KE anisotropy intensity and normalised helicity intensity, in Fig. 7 (a,
 309 b, e, f), we closely examine the KE carried by the moving structural components of the two
 310 BioAVs.

311 As demonstrated in Fig. 7 (a) and (b), the standard deviation of leaflet displacement mag-
 312 nitude over time in the VLth30 case is four times that in the Ulth0 case, as previously
 313 highlighted in the first part of the study⁹. The standard deviation of the displacement mag-
 314 nitude in the supporting ring of the BioAVs is highest at the extremity of the three posts.
 315 In the initial part of the study⁹, hairpin-like vortices were observed near the three posts as a
 316 consequence of the motion of the leaflets and posts, particularly when the gap between the
 317 leaflets and the ring post widens.

318 In Fig. 7 (b), the spatial distribution of the standard deviation of biomechanical KE anisotropy
 319 for the VLth30 case shows strong variation amongst the three leaflets while for the Ulth0
 320 BioAV, $\sigma \left(\mathbb{I}_{p_s}^{\text{flow anis}} \right)_{\text{norm}}$ is more uniformly distributed. With a view to establishing a correla-
 321 tion between the flow and structural anisotropy intensity, we have plotted the graphs depict-
 322 ing the standard deviation of flow modal KE anisotropy and biomechanical KE anisotropy.
 323 These values are averaged over the elevation angles φ and normalised to range from 0 to 1,
 324 as shown in Fig. 7 (c) and (g). The cross-correlation function of the curves in the graphs of
 325 Fig. 7 (c) and (g) has been calculated and is presented in Fig. 7 (d) and (h). It is worth not-
 326 ing that, in the case of the VLth30 BioAV, the graph indicates a strong correlation between
 327 flow and biomechanical anisotropy, as the cross-correlation function reaches its peak at an
 328 azimuthal shift of 0° . In the case of the Ulth0 BioAV, the cross-correlation function in Fig.7
 329 (h) indicates a relatively strong correlation, with peak values occurring at azimuthal shifts
 330 ranging from -10° to 0° . However, the curves in Fig.7 (g) suggest an inverse correlation, as
 331 the minima in $\sigma \left(\mathbb{I}_{p_s}^{\text{flow anis}} \right)_{\text{norm}}$ align with the maxima in $\sigma \left(\mathbb{I}_{p_m}^{\text{struct anis}} \right)_{\text{norm}}$.

332 [Figure 7 about here.]

333 Outlook

334 The findings emphasise the influence of valve architecture on turbulence characteristics in
 335 the ascending aorta near the valvular orifice, with aortic stenosis serving as the benchmark
 336 worst-case scenario. The complex interplay among energy, helicity, leaflet motion and KE
 337 anisotropy is discussed, underscoring the significance of the present work in evaluating BioAV
 338 performance and advancing the development of energy-efficient BioAVs.

339 In fact, the investigation of flow disturbances associated with turbulence, conducted in this
 340 second part of a comprehensive study, relies on advanced energy-based analyses. These
 341 analyses include a dedicated three-dimensional spectral analysis and the introduction of new
 342 quantities, namely the modal and biomechanical KE anisotropy.

343 The detailed two-part computational study lays the foundation for a comprehensive set
 344 of analyses, forming a robust platform for the development of innovative and personalised
 345 valve designs. These designs aim to reduce the adverse consequences linked to aortic valve
 346 replacement.

347 **Methods**

348 The present study constitutes the second part of a comprehensive investigation. Therefore,
 349 for information regarding the geometry of the aortic model, the two valvular bioprostheses
 350 and aortic stenosis along with details about the numerical setups, solving methods and
 351 experimental validation of the solvers, readers are referred to the article that presents the
 352 first part of this extensive study⁹.

353 **Spectral analysis on the blood motion**

354 The spectral analysis proposed in this study relies upon the incompressible Navier-Stokes
 355 equations expressed in Fourier space¹⁴:

$$\boldsymbol{\kappa} \cdot \hat{\mathbf{u}}(\boldsymbol{\kappa}) = 0, \quad (1)$$

356 with $\boldsymbol{\kappa}$, the angular wavenumber vector, \mathbf{u} , the instantaneous flow velocity vector and $\hat{\mathbf{u}}$, the
 357 Fourier modes of \mathbf{u} .

$$\frac{d\hat{\mathbf{u}}(\boldsymbol{\kappa})}{dt} + i \sum_{\boldsymbol{\kappa}'} \{\boldsymbol{\kappa} \cdot \hat{\mathbf{u}}(\boldsymbol{\kappa} - \boldsymbol{\kappa}')\} \hat{\mathbf{u}}(\boldsymbol{\kappa}') = -i\boldsymbol{\kappa} \hat{p}(\boldsymbol{\kappa}) - \nu \kappa^2 \hat{\mathbf{u}}(\boldsymbol{\kappa}), \quad (2)$$

358 with $i = \sqrt{-1}$, ν , the kinematic viscosity and $\hat{p}(\boldsymbol{\kappa})$, the modal kinematic pressure.

359 From these spectral equations, after eliminating the pressure term by projecting the advective
 360 term on the plane of incompressibility¹⁶, an equation for the dynamics of the modal KE
 361 can be derived by taking the inner product of Eq. 2 with the complex-conjugate $\hat{\mathbf{u}}^*(\boldsymbol{\kappa})$
 362 adding the complex-conjugate to the resultant equation. As a result, we obtain the following
 363 equations^{14,23}:

$$\frac{d}{dt} \hat{E}_{u^2}(\boldsymbol{\kappa}) = \underbrace{\sum_{\boldsymbol{\kappa}'} \Im(\{\boldsymbol{\kappa} \cdot \hat{\mathbf{u}}(\boldsymbol{\kappa} - \boldsymbol{\kappa}')\} \{\hat{\mathbf{u}}(\boldsymbol{\kappa}') \cdot \hat{\mathbf{u}}^*(\boldsymbol{\kappa})\})}_{\hat{T}_{u^2}(\boldsymbol{\kappa})} - \underbrace{2\nu\kappa^2 \hat{E}_{u^2}(\boldsymbol{\kappa})}_{\hat{D}_{u^2}(\boldsymbol{\kappa})}, \quad (3)$$

364 with \hat{E}_{u^2} , the modal kinetic energy, \hat{T}_{u^2} , the rate of kinetic energy transfer to the modal
 365 kinetic energy due to non-linearity and \hat{D}_{u^2} , the dissipation rate of modal kinetic energy
 366 through viscous effects. \Im is the imaginary part.

367 It is possible to calculate the total dissipation rate ε by summing $\hat{D}_{u^2}(\boldsymbol{\kappa})$ over the wavenum-
 368 ber. This dissipation ε is used to estimate the Kolmogorov microscales²³.

369 Furthermore, it is suggested in¹² that for non-isotropic turbulence, the spatial velocity func-
 370 tion and the spectrum function be expressed in terms of a single scalar distance r and
 371 wavenumber κ , respectively, by taking mean values of the functions over spherical surfaces
 372 given by radius $r = \text{constant}$ in physical space and $\kappa = \text{constant}$ in Fourier space.

373 This way, the kinetic energy of the Fourier modes \hat{E}_{u^2} of the three-dimensional and anisotropic
 374 velocity field obtained from the simulations is calculated by sampling the velocity compo-
 375 nents on points distributed over spherical shells centred around a point on the centreline of
 376 the ascending aorta (see Fig. 8). A sequence of velocity values (\mathbf{u}_s) is then defined over each
 377 radius of the sphere of points. The points on each spherical shell (i.e. at $r = \text{constant}$) are
 378 obtained by distributing them with a constant increment in both azimuthal and elevation
 379 angles of 30 degrees, resulting in a set of 122 equidistant points on the shell, spaced at an
 380 arc length of $\frac{\pi r}{6}$.

381 [Figure 8 about here.]

382 A one-dimensional formulation of the continuous Fourier transform operator on a finite
 383 interval $(0, \mathcal{L})$ allowing to express the Fourier coefficients of the continuous three-dimensional
 384 velocity field $\mathbf{u}(\boldsymbol{x})$ is defined as follows²³:

$$\mathcal{F}_{\kappa}\{\mathbf{u}(r)\} = \hat{\mathbf{u}}(\kappa) = \frac{1}{\mathcal{L}} \int_0^{\mathcal{L}} \mathbf{u}(r) \exp(-i\kappa r) dr, \quad (4)$$

385 with r , the radial coordinate along each radius of the sphere of points; \mathcal{L} , the radius of
 386 the largest spherical surface used for the sampling. The largest radius is equal to 10 mm
 387 and 11.5 mm for the aortic stenosis and bioprosthesis cases, respectively. It is noteworthy
 388 that the three-dimensional formulation of the Fourier operator is reduced to the above one-
 389 dimensional formulation by integrating along each radius of the sphere of points and by
 390 considering $\boldsymbol{\kappa} \cdot \boldsymbol{x} = \kappa r$ ²³. It is also important to note that the calculation of Fourier modes
 391 implies the periodicity in the velocity field $\mathbf{u}(r)$. However, if we consider the integral length
 392 scale \mathfrak{L} , the effects of this artificially imposed periodicity vanish as $\frac{\mathcal{L}}{\mathfrak{L}}$ tends to infinity²³.
 393 The corresponding discrete Fourier transform (DFT) using a fast Fourier transform (FFT)
 394 algorithm¹¹ is performed based on the instantaneous velocity field \mathbf{u} sampled on a sequence
 395 of points distributed over spherical shells (\mathbf{u}_s) as shown in Fig. 8. The centre of the spherical
 396 shells is positioned on the centreline of the straight aorta at a distance of 12.5 mm and 8
 397 mm from the sino-tubular junction in the stenotic and BioAV cases, respectively. The DFT
 398 is then expressed as:

$$\hat{\mathbf{u}}_k = \frac{1}{(N-1)\Delta r} \sum_{s=0}^{N-1} \mathbf{u}_s \exp\left(\frac{-i2\pi ks}{N}\right) \Delta r, \quad (5)$$

399 with $k = 0, \dots, N-1$. N is the number of points taken over each radius of the spherical
 400 distribution of points and is equal to 82 and 48 for the stenotic and BioAV cases, respectively.
 401 Δr , is the distance between two consecutive spherical shells and is equal to 125 μm for the
 402 stenosis case and 250 μm for the aortic bioprosthesis cases. Each entry of the sequence of
 403 scalar angular wavenumbers κ corresponding to the sequence ($\hat{\mathbf{u}}_k$) is $\kappa_k = \frac{(k+1)}{N} \frac{2\pi}{\Delta r}$ leading
 404 to $\hat{\mathbf{u}}(\kappa) = \hat{\mathbf{u}}_k$.

405 In order to avoid the presence of aliases in the spectrum obtained out of the FFT operation
 406 and considering the symmetry of the spectrum given the real-valued velocity sequence (\mathbf{u}_s),
 407 the values of $\hat{\mathbf{u}}(\kappa)$ above the folding wavenumber $\kappa_f = \frac{\pi}{\Delta r}$ are discarded. These correspond
 408 to the entries $\hat{\mathbf{u}}_k$ with index $k > \lfloor \frac{N+1}{2} \rfloor = N'$.

409 The normalised and discrete first-order autocorrelation function over each radius of the spher-

410 ical distribution of points is then calculated from the modal velocity vector $\hat{\mathbf{u}}_k(\hat{u}_{xk}, \hat{u}_{yk}, \hat{u}_{zk})$:

$$\hat{\mathcal{R}}_I^{u_j u_j} = \frac{1}{|\hat{\mathcal{R}}_0^{u_j u_j}|} \sum_{n=0}^{N'-m-1} (\hat{u}_{j, n+I} \hat{u}_{j, n}^*), \quad (6)$$

411 with $j = (x, y, z)$, $m = 1, \dots, 2N' - 1$, the index $I \in [-N', N']$ and the range of indices I
412 considered for $\hat{\mathcal{R}}^{u_i u_i}$ is $[0, N']$.

413 By inspecting the distribution of the correlation function over the points of each spherical
414 shell, we note a log-normal distribution. Therefore, in order to have the most representative
415 value of the autocorrelation function $\hat{\mathcal{R}}^{u_j u_j}$ over each spherical surface, the mode of this
416 distribution instead of the previously mentioned expected value¹² has been employed to
417 compute an equivalent autocorrelation function $\hat{\mathcal{R}}^{u_j u_j}$ so that the latter is dependent on
418 a single scalar wavenumber κ for each spherical shell. The equivalent correlation function
419 (coming from the mode of the log-normal distribution) is then defined as:

$$\hat{\mathcal{R}}^{u_j u_j} \Big|_{\text{eq}} = \exp(M^{\mathcal{R}} - \Sigma^{\mathcal{R}}), \quad (7)$$

420 $M^{\mathcal{R}} = \log \left[\frac{\langle \hat{\mathcal{R}}^{u_j u_j} \Big|_{\text{sph}} \rangle}{\sqrt{\langle \hat{\mathcal{R}}^{u_j u_j} \Big|_{\text{sph}} \rangle^2 + \mu_2(\hat{\mathcal{R}}^{u_j u_j} \Big|_{\text{sph}})}} \right]$, $\Sigma^{\mathcal{R}} = \log \left[1 + \frac{\mu_2(\hat{\mathcal{R}}^{u_j u_j} \Big|_{\text{sph}})}{\langle \hat{\mathcal{R}}^{u_j u_j} \Big|_{\text{sph}} \rangle^2} \right]$, with $\langle \hat{\mathcal{R}}^{u_j u_j} \Big|_{\text{sph}} \rangle$; the ex-
421 pected value and $\mu_2(\hat{\mathcal{R}}^{u_j u_j} \Big|_{\text{sph}})$; the variance of $\hat{\mathcal{R}}^{u_j u_j}$ over the 122 points of each spherical
422 shell.

423 Finally, each term of Eq. 3 are calculated based on the one-dimensional DFT of the
424 velocity field sampled on points distributed over spherical shells and based on the ensuing
425 equivalent autocorrelation function $\hat{\mathcal{R}}^{u_j u_j} \Big|_{\text{eq}}$. Therefore, the equivalent modal KE $\hat{E}_{u^2}(\kappa)$
426 reads:

$$\hat{E}_{u^2}(\kappa) = \frac{1}{2} \left(\hat{\mathcal{R}}^{u_x u_x} \Big|_{\text{eq}} + \hat{\mathcal{R}}^{u_y u_y} \Big|_{\text{eq}} + \hat{\mathcal{R}}^{u_z u_z} \Big|_{\text{eq}} \right). \quad (8)$$

427 The rate of modal KE transfer $\hat{T}_{u^2}(\kappa)$ is computed as follows:

$$\hat{T}_{u^2}(\kappa) = \frac{1}{2} \left(\hat{T}^{u_x^2} \Big|_{\text{eq}} + \hat{T}^{u_y^2} \Big|_{\text{eq}} + \hat{T}^{u_z^2} \Big|_{\text{eq}} \right), \quad (9)$$

428 where

$$\hat{T}^{u_j^2}(\kappa) = \sum_{\kappa'} \Im \left(\{ \kappa \cdot \hat{u}_j(\kappa - \kappa') \} \{ \hat{u}_j(\kappa') \cdot \hat{u}_j^*(\kappa) \} \right), \quad (10)$$

429 and

$$\hat{T}^{u_j^2}|_{\text{eq}} = \exp(M^T - \Sigma^T), \quad (11)$$

430 with $M^T = \log \left[\frac{\langle \hat{T}^{u_j^2} |_{\text{sph}} \rangle}{\sqrt{\langle \hat{T}^{u_j^2} |_{\text{sph}} \rangle^2 + \mu_2 \langle \hat{T}^{u_j^2} |_{\text{sph}} \rangle}} \right]$, $\Sigma^T = \log \left[1 + \frac{\mu_2 \langle \hat{T}^{u_j^2} |_{\text{sph}} \rangle}{\langle \hat{T}^{u_j^2} |_{\text{sph}} \rangle^2} \right]$.

431 Kinetic energy, enstrophy and helicity in blood flow

432 In order to obtain the fluctuations in the velocity \mathbf{u}' and vorticity $\boldsymbol{\omega}'$ fields of blood flow, a
 433 Reynolds decomposition is performed. With a view of removing the effect of the temporal
 434 periodicity in the leaflet motion on the downstream flow, the time-averaged (over all the con-
 435 sidered time-steps) velocity and vorticity fields are combined with the velocity and vorticity
 436 field phase-averaged at the main frequencies extracted from the leaflet motion analysis⁹.
 437 This decomposition is akin to performing a triple decomposition¹⁵ and leads to the following
 438 formulation for the fluctuating vector fields:

$$\mathbf{u}' = \mathbf{u} - \frac{1}{2}\mathbf{U} - \frac{1}{2}\mathbf{U}_{\text{periodic}} = \mathbf{u} - \underbrace{\frac{1}{2} \left[\frac{1}{T} \int_0^T \mathbf{u} dt + \frac{1}{N} \sum_N \mathbf{u}(t + \frac{N}{f_{\text{mech}}}) \right]}_{\mathbf{U}^{\text{tot}}}, \quad (12)$$

439 with \mathbf{U} , the velocity vector field averaged over the whole simulated systole T ; $\mathbf{U}_{\text{periodic}}$,
 440 the phase-averaged velocity field at frequencies f_{mech} evaluated from the leaflet mechanics
 441 characterisation⁹. The average between these two time-averaged velocity fields gives \mathbf{U}^{tot} .
 442 Likewise, the fluctuating vorticity field is given by:

$$\boldsymbol{\omega}' = \boldsymbol{\omega} - \frac{1}{2}\boldsymbol{\Omega} - \frac{1}{2}\boldsymbol{\Omega}_{\text{periodic}} = \boldsymbol{\omega} - \underbrace{\frac{1}{2} \left[\frac{1}{T} \int_0^T \boldsymbol{\omega} dt + \frac{1}{N} \sum_N \boldsymbol{\omega}(t + \frac{N}{f_{\text{mech}}}) \right]}_{\boldsymbol{\Omega}^{\text{tot}}}, \quad (13)$$

443 with $\boldsymbol{\omega} = \nabla \times \mathbf{u}$, the instantaneous vorticity field; $\boldsymbol{\Omega}$, the mean vorticity vector field time-
 444 averaged over the whole systole, $\boldsymbol{\Omega}_{\text{periodic}}$, the vorticity field phase-averaged at frequencies
 445 evaluated from the leaflet mechanics characterisation⁹ and $\boldsymbol{\Omega}^{\text{tot}}$, the arithmetic average of
 446 the latter two vorticity fields averaged in time.

447 The terms accounting for the periodicity in the flow field due to the periodic leaflet motion
 448 ($\mathbf{u}_{\text{periodic}}$ and $\boldsymbol{\omega}_{\text{periodic}}$) are obviously null in the stenotic case and in the case where the leaflets
 449 are not moving.

450 The total turbulent KE, fluctuating enstrophy and fluctuating helicity are given by:

$$TE_{u'^2} = \int_r \frac{1}{2} (\mathbf{u}'(r) \cdot \mathbf{u}'(r)) dr, \quad (14)$$

$$TE_{\omega'^2} = \int_r \frac{1}{2} (\boldsymbol{\omega}'(r) \cdot \boldsymbol{\omega}'(r)) dr, \quad (15)$$

$$TE_{|h'|} = \int_r \frac{1}{2} |\mathbf{u}'(r) \cdot \boldsymbol{\omega}'(r)| dr, \quad (16)$$

$$TE_{h'} = \int_r \frac{1}{2} (\mathbf{u}'(r) \cdot \boldsymbol{\omega}'(r)) dr, \quad (17)$$

451 with r , the coordinate along each radius corresponding to the radius from the centre point
 452 of the spherical distribution of points defined in the spectral analysis section.

453 The mean KE, enstrophy and helicity are defined as:

$$TE_{U^2} = \int_r \frac{1}{2} (\mathbf{U}^{\text{tot}}(r) \cdot \mathbf{U}^{\text{tot}}(r)) dr, \quad (18)$$

$$TE_{\Omega^2} = \int_r \frac{1}{2} (\boldsymbol{\Omega}^{\text{tot}}(r) \cdot \boldsymbol{\Omega}^{\text{tot}}(r)) dr, \quad (19)$$

$$TE_{|H|} = \int_r \frac{1}{2} |\mathbf{U}^{\text{tot}}(r) \cdot \boldsymbol{\Omega}^{\text{tot}}(r)| dr, \quad (20)$$

$$TE_H = \int_r \frac{1}{2} (\mathbf{U}^{\text{tot}}(r) \cdot \boldsymbol{\Omega}^{\text{tot}}(r)) dr. \quad (21)$$

454 The turbulence intensity $\mathbb{I}_{u'^2}$ (Eq. 22), intensity of fluctuating enstrophy $\mathbb{I}_{\omega'^2}$ (Eq. 23), inten-
 455 sity of unsigned fluctuating helicity $\mathbb{I}_{|h'|}$ (Eq. 24) and intensity of signed fluctuating helicity
 456 $\mathbb{I}_{h'}$ (Eq. 25) are calculated by taking the ratio of the aforementioned fluctuating and mean
 457 quantity fields:

$$\mathbb{I}_{u'^2} = \frac{TE_{u'^2}}{TE_U^2} \quad (22)$$

$$\mathbb{I}_{\omega'^2} = \frac{TE_{\omega'^2}}{TE_{\Omega^2}} \quad (23)$$

$$\mathbb{I}_{|h'|} = \frac{TE_{|h'|}}{TE_{|H|}} \quad (24)$$

$$\mathbb{I}_{h'} = \frac{TE_{h'}}{TE_H} \quad (25)$$

458 Akin to the calculation of the equivalent autocorrelation function $\hat{\mathcal{R}}^{u_j u_j}$ and the correspond-
 459 ing modal KE \hat{E}_{u^2} , we compute an equivalent value for the foregoing intensity fields at each
 460 time instance under consideration. This equivalent value represents the most representative
 461 intensity field value over the 122 points of the spherical shells and it is determined by taking
 462 the mode of the log-normal distribution (cf. Eq. 7).

463 Flow modal kinetic energy and helicity intensity

464 The total flow KE \mathcal{K}_{p_s} for each point p_s of the spherical shell is calculated from the normalised
 465 and discrete first-order autocorrelation function as follows:

$$\mathcal{K}_{p_s} = \frac{1}{2} \sum_j \int_{\kappa} \hat{\mathcal{R}}^{u_j u_j} |_{\text{sph}} d\kappa, \quad (26)$$

466 with $j = (x, y, z)$ and $p_s = 1, \dots, 122$, which corresponds to the index of each point on the
 467 spherical shell. The numerical integration is performed using the trapezoidal rule.

468 The flow modal KE anisotropy $\mathbb{I}_{p_s}^{\text{flow anis}}$ over a spherical region close to the aortic orifice is
 469 thus defined as:

$$\mathbb{I}_{p_s}^{\text{flow anis}} = \frac{\mathcal{K}_{p_s} - \langle \mathcal{K}_{p_s} \rangle}{\langle \mathcal{K}_{p_s} \rangle} \times 100, \quad (27)$$

470 with $\langle \cdot \rangle$, the average operator over the 122 points of the spherical shell.

471 Similarly, a normalised helicity intensity $\mathbb{I}_{p_s}^{h|\text{sph}}$ is computed along each radial direction of the
 472 spherical shells from the instantaneous helicity intensity field through the following equation:

$$\frac{\mathbb{I}_{p_s}^{h|\text{sph}}}{100} = \frac{\int_r \frac{1}{2} (\mathbf{u}(r) \cdot \boldsymbol{\omega}(r)) dr}{\langle \int_r \frac{1}{2} (\mathbf{u}(r) \cdot \boldsymbol{\omega}(r)) dr \rangle} = \frac{TE_h|_{\text{sph}}}{\langle TE_h|_{\text{sph}} \rangle} = \frac{\mathbb{I}_h|_{\text{sph}}}{\langle \mathbb{I}_h|_{\text{sph}} \rangle}. \quad (28)$$

473 **Anisotropy in leaflet and ring kinetic energy**

474 The KE computed at the mesh points p_m located at the leaflet or ring interfaces of the
 475 simulated bioprosthesis valve motion is expressed as:

$$KE_{p_m}^{\text{struct}} = \frac{1}{2} \sum_j \mathbf{v}_j^2|_{\text{int}}, \quad (29)$$

476 with $\mathbf{v}_j|_{\text{int}}$, the j^{th} velocity component of the structure at the fluid-solid interface (int), which
 477 is equal to the flow velocity at the interface by virtue of the velocity continuity condition.

478 The KE anisotropy $\mathbb{I}_{p_m}^{\text{struct anis}}$ in the structure (ring and leaflet) is then defined as:

$$\frac{\mathbb{I}_{p_m}^{\text{struct anis}}}{100} = \left[\frac{KE_{p_m}^{\text{struct}} - \langle KE_{p_m}^{\text{struct}} \rangle}{\langle KE_{p_m}^{\text{struct}} \rangle} \right] (\mathbf{p}^{\text{struct}} \cdot \mathbf{n}^{\text{struct}}), \quad (30)$$

479 with $\mathbf{p}^{\text{struct}}$, the coordinates of mesh points of the considered structure (leaflet or ring) and
 480 $\mathbf{n}^{\text{struct}}$, the outward normal vector to the considered mesh point of the structure. The operator
 481 $\langle \cdot \rangle$ corresponds to the average over the different mesh points of the structure.

482 **Acknowledgements**

483 This work was supported by the computational resources from the Swiss National Super-
 484 computing Centre (CSCS) under project IDs c12, s1012 and sm56.

485 **Author contribution**

486 **P. Corso:** Conceptualisation, Data curation, Formal analysis, Interpretation of the results,
487 Investigation, Methodology, Software, Visualisation, Writing - original draft. **D. Obrist:**
488 Funding acquisition, Input on the results and on the original draft.

489 **Declaration of competing interest**

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

492 **Data and materials availability**

493 All data and materials needed to evaluate the conclusions of this paper are present in the
494 main text or supplementary materials. Processable data files can be obtained from the first
495 author.

496 **References**

- 497 1. Aluru J. S., Barsouk A., Saginala K., Rawla P., Barsouk A., *Valvular Heart Disease*
498 *Epidemiology, Medical Sciences*, 10, 2022.
- 499 2. Bailoor S., Seo J.-H., Dasi L., *et al.* *A computational study of the hemodynamics of*
500 *bioprosthetic aortic valves with reduced leaflet motion*, *J. of Biomechanics*, 120, 2021.
- 501 3. Baumgartner H., Falk V., Bax J. J., De Bonis M., Hamm C., Holm P. J., Lung
502 B., Lancellotti P., Lansac E., Rodriguez Muñoz D., Rosenhek R., Sjögren J., Tornos
503 Mas P., Vahanian A., Walther T., Wendler O., Windecker S., Zamorano J. L. 2017
504 ESC/EACTS Guidelines for the management of valvular heart disease. *European Heart*
505 *J.* 38(36):2739–2791, 2017.

- 506 4. Becsek B., Pietrasanta L. and Obrist D., *Turbulent Systolic Flow Downstream of a Bio-*
507 *prosthetic Aortic Valve: Velocity Spectra, Wall Shear Stresses, and Turbulent Dissipation*
508 *Rates*, Front. in physiol. 11:539-575, 2020.
- 509 5. Caballero A.D., Laín S.J.C.E., *A review on computational fluid dynamics modelling in*
510 *human thoracic aorta*, Cardiovasc. Engng Technol., 4(2):103–130, 2013.
- 511 6. Corso P., Gülan U., Cohrs N., *et al.*, *Comprehensive In Vitro Study of the Flow Past Two*
512 *Transcatheter Aortic Valves: Comparison with a Severe Stenotic Case*, Ann. Biomed.
513 Eng., 47(11):2241-2257, 2019.
- 514 7. Corso P., Giannakopoulos G., Gulan U., *et al.*, *A Novel Estimation Approach of Pressure*
515 *Gradient and Haemodynamic Stresses as Indicators of Pathological Aortic Flow Using*
516 *Subvoxel Modelling*, IEEE Trans. Biomed. Eng., 68(3):980-991, 2021.
- 517 8. Corso P., Walheim J., Dillinger H., *et al.*, *Toward an accurate estimation of wall shear*
518 *stress from 4D flow magnetic resonance downstream of a severe stenosis*, Magn. Res.
519 Med. 86(3):1531-1543, 2021.
- 520 9. Corso P., Obrist D., *On the role of aortic valve architecture for physiological haemody-*
521 *namics and valve replacement. Part I: flow topology and vortex dynamics*, 2023.
- 522 10. Davidson P., *Turbulence: An Introduction for Scientists and Engineers - 2nd Edition*,
523 Oxford University Press, 2015.
- 524 11. Frigo M. and Johnson S. G., *The Design and Implementation of FFTW3*, Proceedings
525 of the IEEE, 93(2):216–231, 2005.
- 526 12. Frost W., Moulden T. H., *Handbook of Turbulence: volume 1 - Fundamentals and Ap-*
527 *plications*, Plenum Press, 1977
- 528 13. Gallo D., Morbiducci U., de Tullio M. D., *On the unexplored relationship between kinetic*
529 *energy and helicity in prosthetic heart valves hemodynamics*, Int. J. Eng. Science, 117,
530 2022.

- 531 14. Gupta A., Jayaram R., Chatterjee A. G., *et al.*, *Energy and enstrophy spectra and fluxes*
532 *for the inertial-dissipation range of two-dimensional turbulence*, Physical Review E, 100,
533 2019.
- 534 15. Hussain A. K. M. F., Reynolds W. C., *The mechanics of an organized wave in turbulent*
535 *shear flow*. J. Fluid Mech., 41(2):241-258, 1970.
- 536 16. Lesieur M. *Turbulence in Fluids: Fourier Analysis of Homogeneous Turbulence*, Springer
537 Dordrecht, 3rd Edition, 1997.
- 538 17. Lindman B. R., Clavel M.-A., Mathieu P., Lung B., Lancellotti P., Otto C. M., Pibarot
539 Ph. *Calcific aortic stenosis*. Nature Reviews Disease Primers 2, 2016.
- 540 18. Moffatt, H. K., *The degree of knottedness of tangled vortex lines*. J. Fluid Mech., 36:
541 117-129, 1969.
- 542 19. Moffat H. K., Tsinober A., *Helicity in laminar and turbulent flow*, Ann. Rev. Fluid
543 Mech., 24:281-312, 1992.
- 544 20. Morbiducci U., Ponzini R., Rizzo G., *et al.*, *Mechanistic insight into the physiological*
545 *relevance of helical blood flow in the human aorta: An in vivo study*, Biomechanics and
546 Modeling in Mechanobiology, 10:339-355, 2011.
- 547 21. Nguyen V., Michel M., Eltchaninoff H., *et al.*, *Implementation of Transcatheter Aortic*
548 *Valve Replacement in France*, J. Am. College Card., 71(15), 2018.
- 549 22. Kitware and Inc. *VTK User's Guide* Kitware Inc., 11th edition, 3 2010. ISBN 978-
550 1930934238.
- 551 23. Pope S. B., *Turbulent Flows*, Cambridge University Press, 2000.
- 552 24. Pujari S. H., Agasthi P. Aortic Stenosis. [Updated 2022 Apr 14]. In: StatPearls
553 [Internet]. Treasure Island (FL): StatPearls Publishing; 2022 Jan-. Available from:
554 <https://www.ncbi.nlm.nih.gov/books/NBK557628/>

- 555 25. Sotiropoulos F., Le T. B., Gilmanov, A. *Fluid mechanics of heart valves and their re-*
556 *placements*. Annual Review of Fluid Mechanics 48, 259–283, 2016.
- 557 26. Townsend A. A., *Entrainment and the structure of turbulent flow*. J. Fluid Mech., 41:13-
558 46, 1970.
- 559 27. Tsolaki E., Corso P., Zboray R., *et al.*, *Multiscale multimodal characterization and sim-*
560 *ulation of structural alterations in failed bioprosthetic heart valves*. Acta Biomaterialia,
561 169:138-154, 2023.
- 562 28. Varghese S. S., Frankel S. H., Fischer P., . *Direct numerical simulation of stenotic flows.*
563 *Part 1. Steady flow*, J. Fluid Mech., 582:253-280, 2007.
- 564 29. Yoganathan A. J., He S., Jones S. C. *Fluid mechanics of heart valves*. Annu. Rev. Biomed.
565 Eng.,6:331–362, 2004.

566 List of Figures

567	1	Three-dimensional wavenumber spectra of kinetic energy for the three valvular cases are shown in (a), (b) and (c). These spectra represent the spectral curve for all the considered time instances between $t = 0.1$ and $t = 0.3$ s, highlighting the maximum, minimum, and median values at each scalar wavenumber. (d), (e), and (f) display the derivative of the modal KE with respect to the angular wavenumber to determine whether the well-known $-5/3$ power-law decay is observed in the inertial subrange.	28
568	2	Time series of turbulence characteristics for the three valvular cases over the systolic phase. (a) Kolmogorov length scale. (b) Dissipation rate of turbulent kinetic energy ε . (c) Integral length scale \mathfrak{L}	29
569	3	Time series over systole of normalised (a) intensity of fluctuating KE, (b) intensity of fluctuating enstrophy, (c) intensity of unsigned fluctuating helicity and (d) intensity of signed fluctuating helicity. The three valvular cases investigated in this study are plotted along with the valvular case presented in Gallo <i>et al.</i> ¹³ . (e) Spider chart of the time-averaged ($\bar{}$) and maximum normalised $\mathbb{I}_{u'^2}$ (I and II) and $\mathbb{I}_{ h' ^2}$ (III and IV). The maximum and absolute minimum values of normalised $\mathbb{I}_{h'^2}$ is also displayed (V and VI).	30
570	4	Probability distributions of the modal KE anisotropy intensity. (a) Fitted log-normal probability density function (PDF) for the three valve configurations, the mode of the PDFs was used in the spectral analysis part of the study. The rhombi represent the modes of the log-normal distribution fitted for each instant considered over peak systole. (b) Boxplots and distribution for the comparison of the shape of the distributions for the three valvular cases.	31
571	5	Spatial distribution and statistical description, including the standard deviation σ and the temporal average value of the time-dependent modal KE anisotropy fields $ \mathbb{I}_{p_s}^{\text{flow anis}} $ obtained on spherical shells near the valvular orifice. The geometries of the spheres and orifice are projected onto a rectangular map using a cylindrical map projection, also known as the Mercator projection. In this map, the azimuthal angle α is represented on the x-axis and the elevation angle φ is represented on the y-axis. The stenotic case is presented in (a, b, c), the valvular case with the design VLth30 in (d, e, f) and the case of the BioAV with the design Ulth0 in (g, h, i).	32
572			
573			
574			
575			
576			
577			
578			
579			
580			
581			
582			
583			
584			
585			
586			
587			
588			
589			
590			
591			
592			
593			
594			
595			
596			
597			
598			

599	6	Anti-correlation between $\overline{ \mathbb{I}_{p_s}^{\text{flow anis}} }$ and $\overline{\mathbb{I}_{p_s}^{h \text{sph}}}$ over spherical shells close to the valvular orifice (cf. Fig. 8). The colour-coded distribution map of the temporal average of $ \mathbb{I}_{p_s}^{\text{flow anis}} $ for the three valvular configurations under consideration is displayed in (a, e, i). The distribution heat map of time-averaged normalised helicity intensity $\mathbb{I}_{p_s}^{h \text{sph}}$ is displayed in (b, f, j) for the three valvular cases. (c, g, k) Scatter plots of $\overline{ \mathbb{I}_{p_s}^{\text{flow anis}} }$ as a function of $\overline{\mathbb{I}_{p_s}^{h \text{sph}}}$. The two coefficients A and B of a power law correlating the two quantities are fitted through the resolution of a non-linear least-square minimisation problem ^{7,8,27} . The accuracy of the training prediction is evaluated using the coefficient of determination R^2 . The accuracy of the prediction based on the testing dataset points for the three valvular cases is presented in (d, h, l).	33
610	7	Comparison and correlation between the anisotropy in the leaflet and ring motion of the two bioprosthetic valves (VLth30 and Ulth0) and anisotropy of the modal kinetic energy in the flow in the vicinity of the valve orifice. (a, e) Standard deviation σ of the unrolled point distribution of the anisotropy intensity based upon the valve kinetic energy. (b, f) Standard deviation σ of the time-dependent unrolled spatial distribution of the displacement magnitude in the two valves. (c, g) Comparison of the standard deviation of the anisotropy intensity in the valve motion and in the flow averaged over the elevation angle φ as a function of the angle α . (g, h) Cross-correlation functions between the two $\sigma(\mathbb{I}_{p_s}^{\text{flow anis}})_{\text{norm}}$ and $\sigma(\mathbb{I}_{p_m}^{\text{struct anis}})_{\text{norm}}$ curves as a function of α . 34	34
620	8	Definition of the points on spherical shells for the computation of the Fourier modes and of the first-order autocorrelation function as part of the spectral analysis for (a) the stenotic case and (b) one of the BioAV cases (VLth30). The spherical surface consists of 122 points as displayed in (c) equally spaced by a distance Δr of 125 m in the stenotic case and 250 m in the BioAV cases. The largest radius \mathcal{L} for the spherical shell is equal to 10 mm and 11.5 mm for the stenotic and BioAV cases, respectively. The centre of the spheres is on the centreline of the straight ascending aorta. The instantaneous velocity magnitude is visualised in (a) and (b) through volumetric rendering displaying the jet of high velocity issuing from the valvular orifice.	35

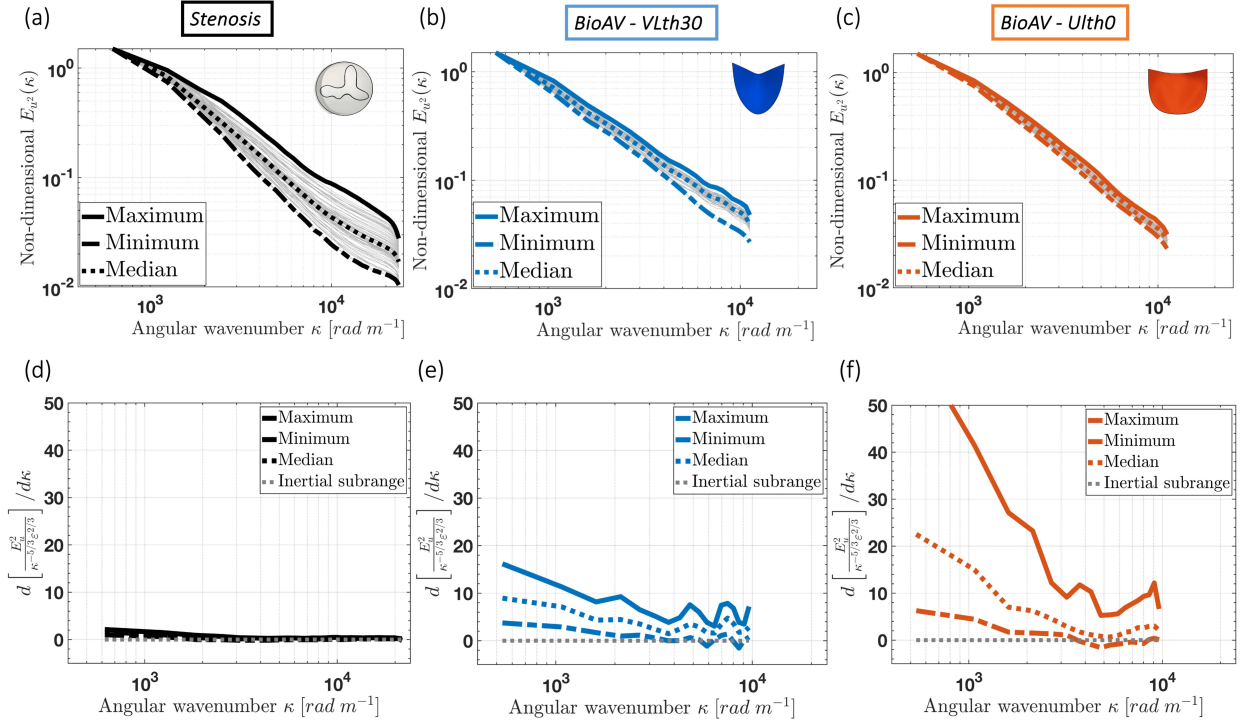


Figure 1: Three-dimensional wavenumber spectra of kinetic energy for the three valvular cases are shown in (a), (b) and (c). These spectra represent the spectral curve for all the considered time instances between $t = 0.1$ and $t = 0.3$ s, highlighting the maximum, minimum, and median values at each scalar wavenumber. (d), (e), and (f) display the derivative of the modal KE with respect to the angular wavenumber to determine whether the well-known $-5/3$ power-law decay is observed in the inertial subrange.

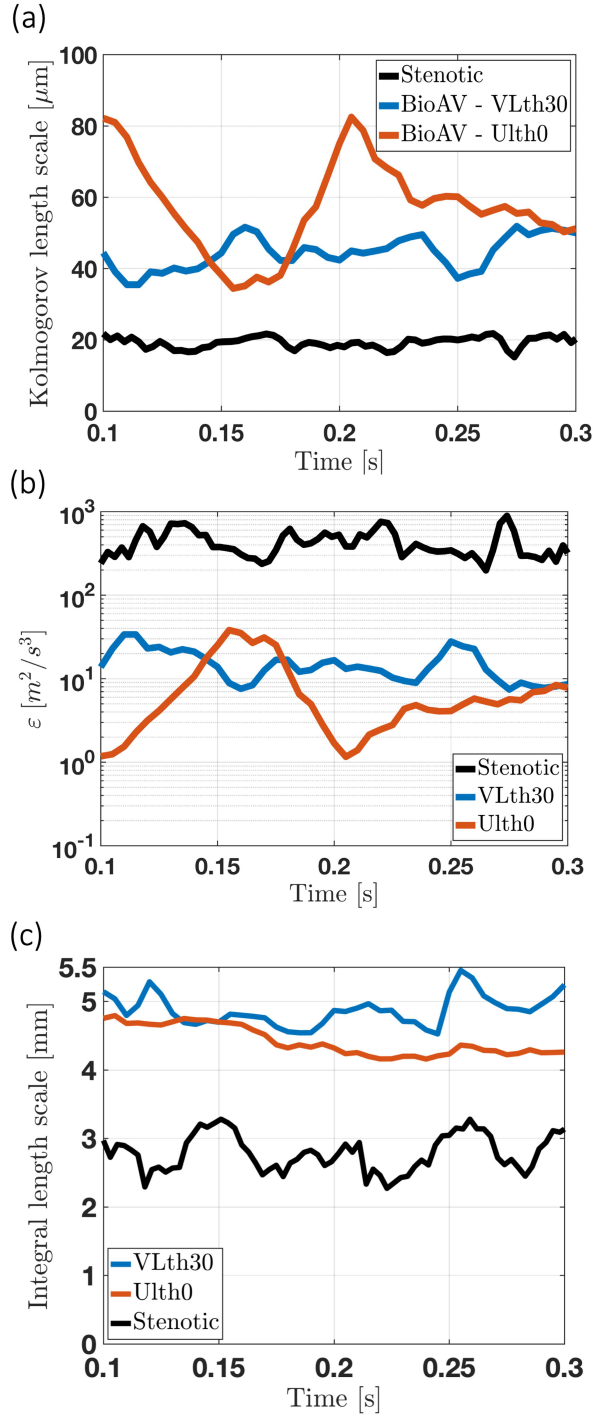


Figure 2: Time series of turbulence characteristics for the three valvular cases over the systolic phase. (a) Kolmogorov length scale. (b) Dissipation rate of turbulent kinetic energy ε . (c) Integral length scale \mathcal{L} .

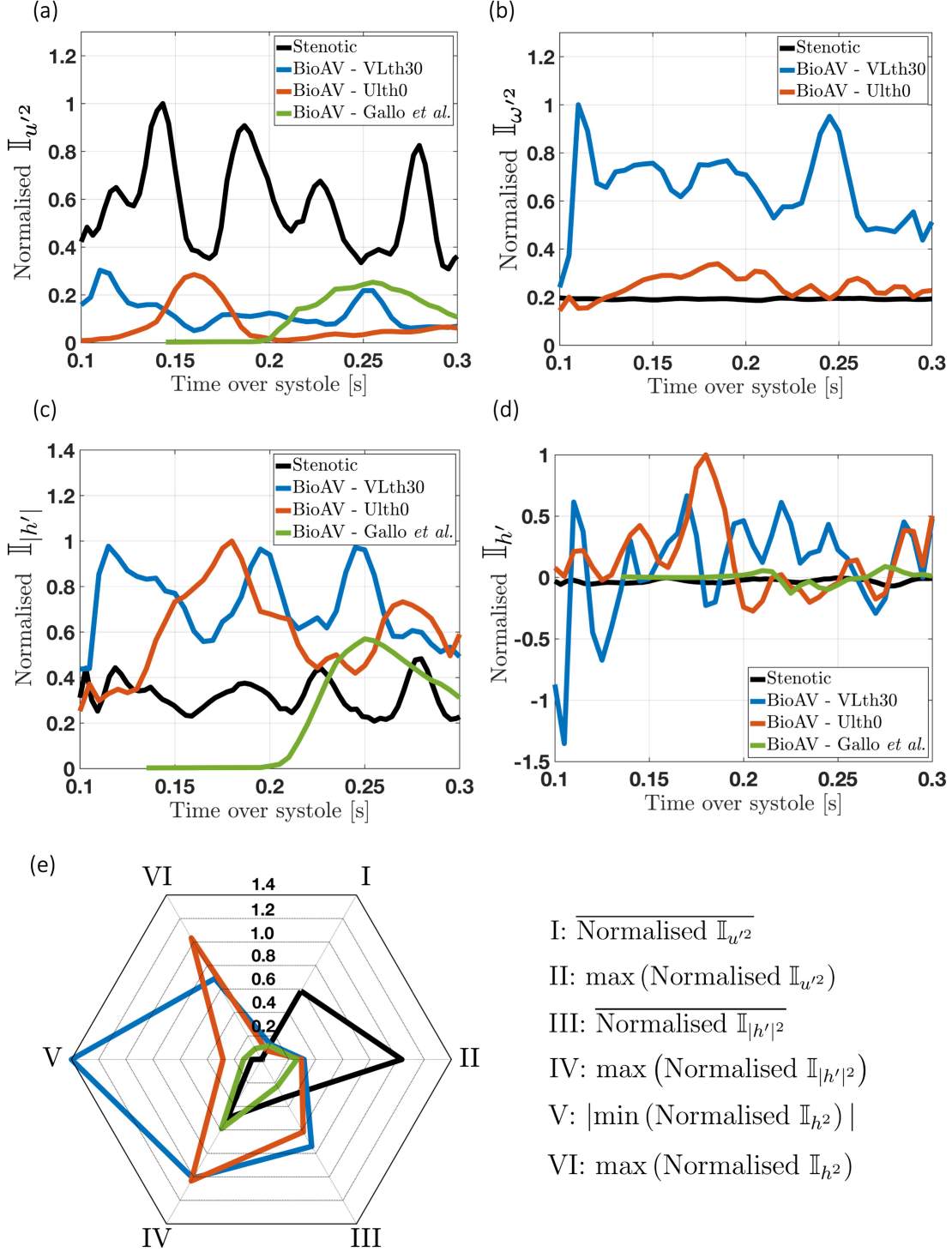


Figure 3: Time series over systole of normalised (a) intensity of fluctuating KE, (b) intensity of fluctuating enstrophy, (c) intensity of unsigned fluctuating helicity and (d) intensity of signed fluctuating helicity. The three valvular cases investigated in this study are plotted along with the valvular case presented in Gallo *et al.*¹³. (e) Spider chart of the time-averaged ($\bar{\cdot}$) and maximum normalised $\mathbb{I}_{u'^2}$ (I and II) and $\mathbb{I}_{|h'|^2}$ (III and IV). The maximum and absolute minimum values of normalised $\mathbb{I}_{h'^2}$ is also displayed (V and VI).

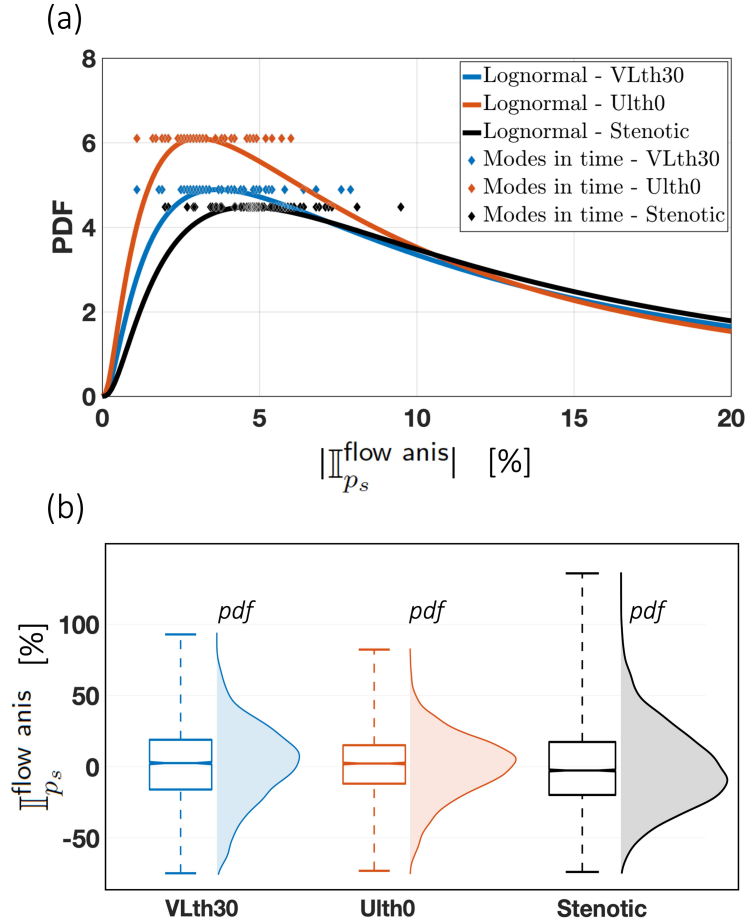


Figure 4: Probability distributions of the modal KE anisotropy intensity. (a) Fitted log-normal probability density function (PDF) for the three valve configurations, the mode of the PDFs was used in the spectral analysis part of the study. The rhombi represent the modes of the log-normal distribution fitted for each instant considered over peak systole. (b) Boxplots and distribution for the comparison of the shape of the distributions for the three valvular cases.

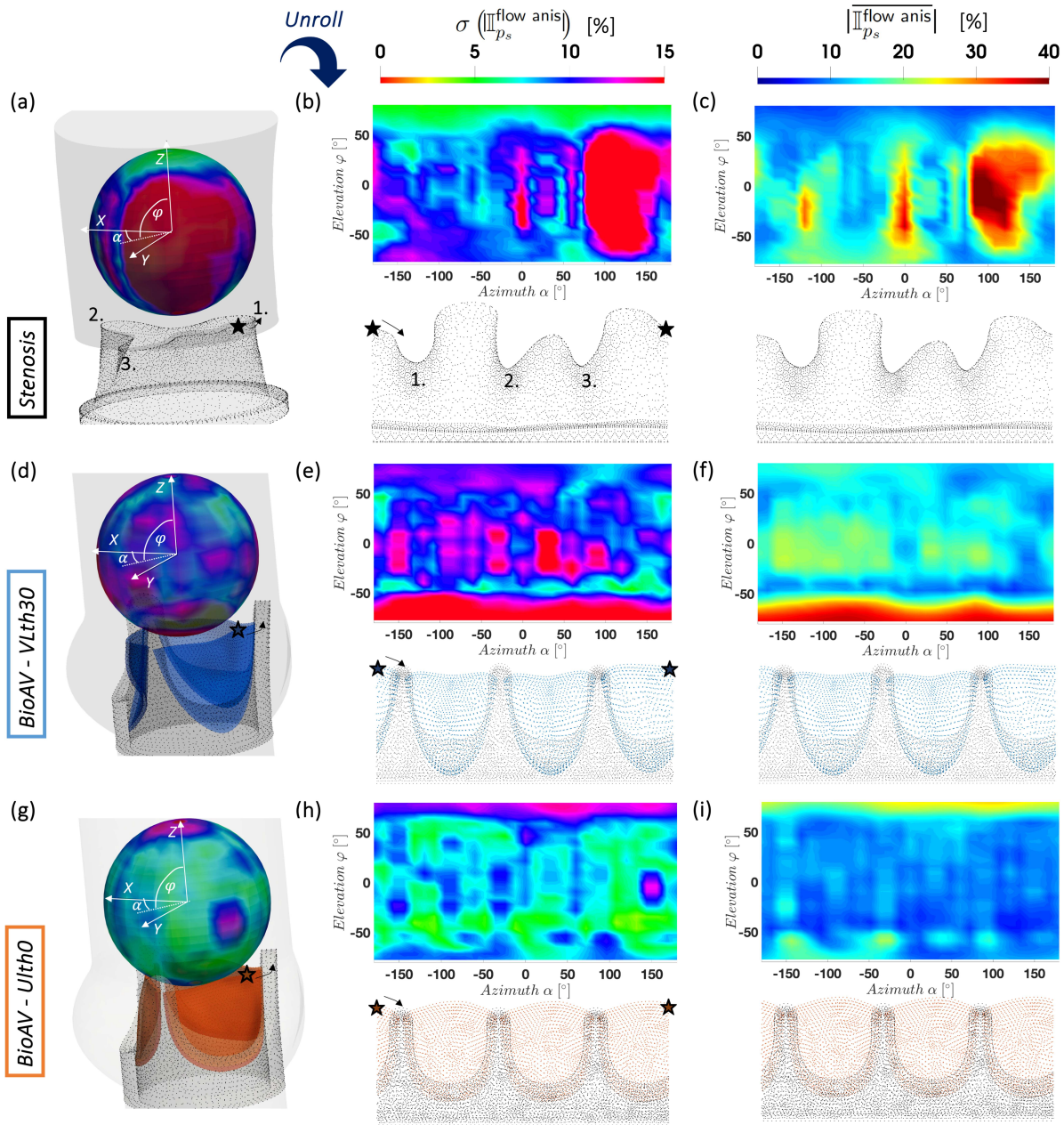


Figure 5: Spatial distribution and statistical description, including the standard deviation σ and the temporal average value of the time-dependent modal KE anisotropy fields $|\overline{II}_{p_s}^{\text{flow anis}}|$ obtained on spherical shells near the valvular orifice. The geometries of the spheres and orifice are projected onto a rectangular map using a cylindrical map projection, also known as the Mercator projection. In this map, the azimuthal angle α is represented on the x-axis and the elevation angle φ is represented on the y-axis. The stenotic case is presented in (a, b, c), the valvular case with the design VLth30 in (d, e, f) and the case of the BioAV with the design Ulth0 in (g, h, i).

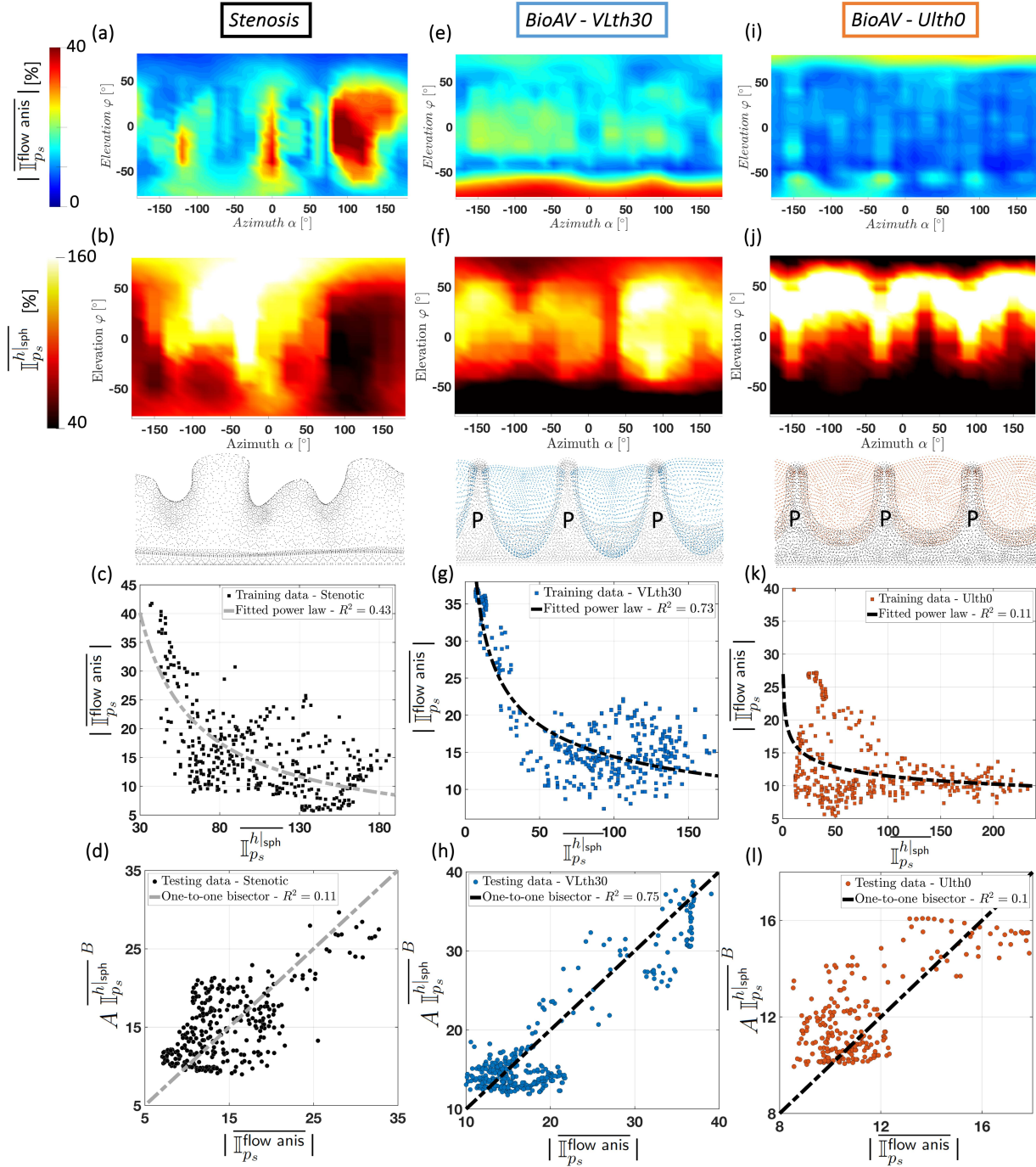


Figure 6: Anti-correlation between $\overline{|\mathbb{I}_{p_s}^{\text{flow anis}}|}$ and $\overline{\mathbb{I}_{p_s}^{h|\text{sph}}}$ over spherical shells close to the valvular orifice (cf. Fig. 8). The colour-coded distribution map of the temporal average of $|\mathbb{I}_{p_s}^{\text{flow anis}}|$ for the three valvular configurations under consideration is displayed in (a, e, i). The distribution heat map of time-averaged normalised helicity intensity $\mathbb{I}_{p_s}^{h|\text{sph}}$ is displayed in (b, f, j) for the three valvular cases. (c, g, k) Scatter plots of $\overline{|\mathbb{I}_{p_s}^{\text{flow anis}}|}$ as a function of $\overline{\mathbb{I}_{p_s}^{h|\text{sph}}}$. The two coefficients A and B of a power law correlating the two quantities are fitted through the resolution of a non-linear least-square minimisation problem^{7,8,27}. The accuracy of the training prediction is evaluated using the coefficient of determination R^2 . The accuracy of the prediction based on the testing dataset points for the three valvular cases is presented in (d, h, l).

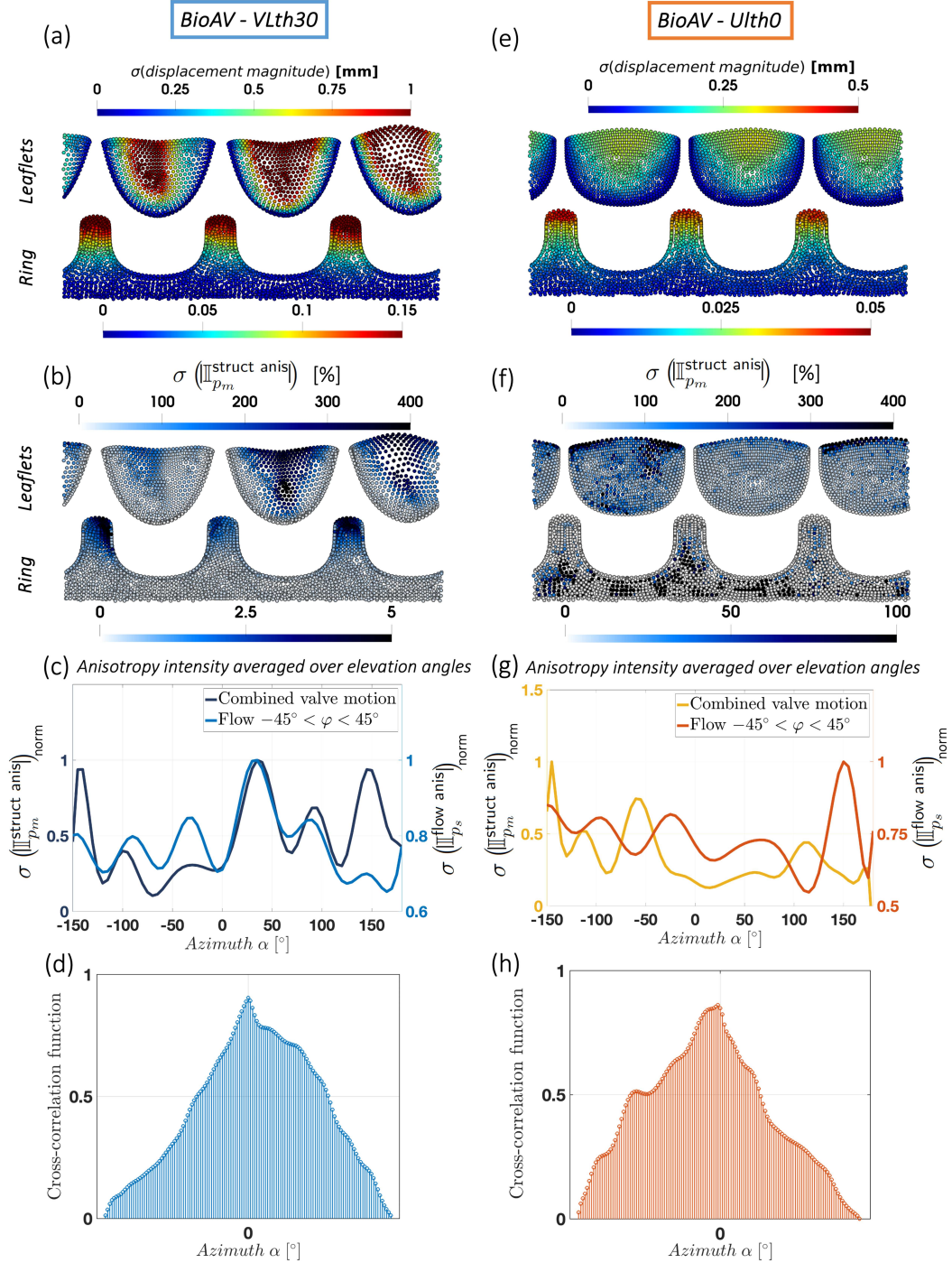


Figure 7: Comparison and correlation between the anisotropy in the leaflet and ring motion of the two bioprosthetic valves (VLth30 and Ulth0) and anisotropy of the modal kinetic energy in the flow in the vicinity of the valve orifice. (a, e) Standard deviation σ of the unrolled point distribution of the anisotropy intensity based upon the valve kinetic energy. (b, f) Standard deviation σ of the time-dependent unrolled spatial distribution of the displacement magnitude in the two valves. (c, g) Comparison of the standard deviation of the anisotropy intensity in the valve motion and in the flow averaged over the elevation angle φ as a function of the angle α . (g, h) Cross-correlation functions between the two σ (II_{p_s}^{flow anis})_{norm} and σ (II_{p_m}^{struct anis})_{norm} curves as a function of α .

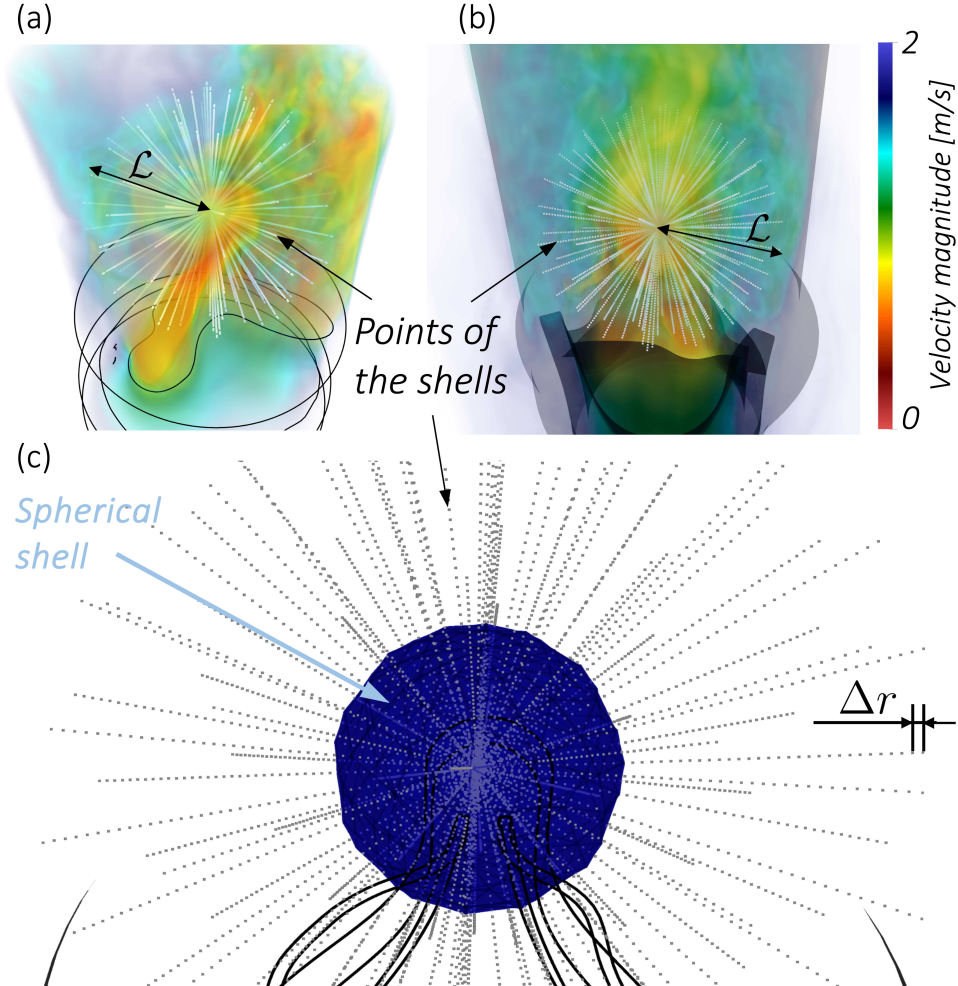


Figure 8: Definition of the points on spherical shells for the computation of the Fourier modes and of the first-order autocorrelation function as part of the spectral analysis for (a) the stenotic case and (b) one of the BioAV cases (VLth30). The spherical surface consists of 122 points as displayed in (c) equally spaced by a distance Δr of 125 m in the stenotic case and 250 m in the BioAV cases. The largest radius \mathcal{L} for the spherical shell is equal to 10 mm and 11.5 mm for the stenotic and BioAV cases, respectively. The centre of the spheres is on the centreline of the straight ascending aorta. The instantaneous velocity magnitude is visualised in (a) and (b) through volumetric rendering displaying the jet of high velocity issuing from the valvular orifice.

Supplementary Files

This is a list of supplementary files associated with this preprint. Click to download.

- [ManuscriptvalveDesignPart2PasSINatBiomed.pdf](#)