been followed to guarantee impartial management of the manuscript.

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REFERENCES

- 1. Garcia-Pavia P, Domínguez F, Gonzalez-Lopez E. Transthyretin amyloid cardiomyopathy. Med Clin (Barc). 2021;156:126-134.
- 2. Álvarez Rubio J, Manovel Sánchez AJ, González-Costello J, et al. Characterization of hereditary transthyretin cardiac amyloidosis in Spain. Rev Esp Cardiol. 2022:75:488-495
- 3. Garcia-Pavia P, Rapezzi C, Adler Y, et al.Diagnosis and treatment of cardiac amyloidosis: a position statement of the ESC Working Group on Myocardial and Pericardial Diseases. Eur Heart J. 2021;42:1554-1568.
- 4. Daoko J, Elnahar Y, el Kersh K, Mohammad N, Shamoon F, Cardiac MRI. detection of a rare case of familial cardiac amyloidosis (Ser23asn): Case report with literature review. Rep Med Imaging. 2010;3:123-127.
- 5. Castaño A, Bokhari S, Brannagan TH, Wynn J, Maurer MS. Technetium pyrophosphate myocardial uptake and peripheral neuropathy in a rare variant of familial transthyretin (TTR) amyloidosis (Ser23Asn): A case report and literature review. Amyloid, 2012:19:41-46.
- 6. Papathanasiou M, Carpinteiro A, Kersting D, et al. Rare variant (p.Ser43Asn) of familial transthyretin amyloidosis associated with isolated cardiac phenotype: A case series with literature review. Mol Genet Genomic Med. 2021. http://dx.doi.org/ 10.1002/mgg3.1581.

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Fluid-dynamic impact of commissural alignment in transcatheter aortic valve implantation

Impacto fluidodinámico del alineamiento comisural en implante percutáneo de válvula aórtica

To the Editor,

New commissural alignment techniques in transcatheter aortic valve implantation (TAVI) can facilitate coronary reaccess, if needed, and prevent coronary obstruction in valve-in-valve TAVI procedures. These techniques could also reduce central aortic regurgitation, gradient progression, and subclinical thrombosis, which would improve durability, all of which are crucial in low-risk patients. However, in specific patients it is impossible to determine differences between an implant with correct neocommissural alignment and one with misalignment. Computational fluid dynamics models have been widely validated for the cardiovascular system¹ and could help to predict the impact of different degrees of commissural misalignment on transvalvular gradients² and the degree of platelet activation in specific patients.³ Based on computed tomography (CT) scans of the patients included in this study, we estimated the central axis of the aorta and predicted the degree of clockwise/counterclockwise rotation required by the delivery system (self-expanding) or the prosthesis at crimping (balloon expandable) for correct commissural alignment using a previously described technique.⁴ After implantation, CT/angiography was used to assess the degree of alignment. Based on the patients' anatomical and pressure data, computational fluid dynamics simulations were performed using the calculated velocity fields (figure 1A; video 1 of the supplementary data), with the valve fully open, by

parametric study of the influence of the commissural alignment angle, rotating the valve in incremental 1-degree steps from perfect alignment (0°) up to 119° . These simulations were performed for two situations: *a*) an unrealistic completely uniform flow at the ventricular outflow tract, as simulated in current models (figure 1C); and (b) a helical flow similar to that used in MRI studies (figure 1D). The platelet activation model used was the same as the one validated in an in-vitro model of coronary bifurcation.³

Forty patients with severe aortic stenosis, trileaflet aortic stenosis, severe calcification, and an aortic annulus area of $478 \pm 101 \text{ mm}^2$ (age, 81.4 ± 4.9 years; STS risk score, 5.2 ± 2.9) were treated with TAVI with commissural alignment. Ventricular function was $57.5\pm11.9\%$ and mean gradient was 51.1 ± 18.5 mmHg, increasing to 6.3 ± 2.1 mmHg at 30 days, with no differences related to alignment (0% mortality). Mean commissural misalignment was 9.2° (ie, moderate; $< 30^{\circ}$) in all patients. Significant differences in alignment were found between selfexpanding prostheses (Acurate neo [Boston Scientific, EE. UU.] and Evolut [Medtronic, EE. UU.]: 7 \pm 3.2°; n = 28) and balloon expandable prostheses (SAPIEN-3/Ultra [Edwards, EE. UU.] and MyVal [Meril, India]: $11.2^{\circ} \pm 4.9^{\circ}$; *P* = .001; n = 12). Analysis of the 2 fluid dynamics simulations, as described above, (figure 2A), showed high sensitivity to ventricular outflow conditions, such that in the more realistic (helical) flow model an association was found between the absence of moderate or severe commissural misalignment (around 30° or less) and higher ventricular energy efficiency (lower line, red). However, in the uniform flow model, no association was found between better commissural alignment and reduced shear stress (upper line, blue). No differences were found between the 2 conditions (helical or uniform) in the degree of platelet activation according to previously validated models (figure 2B).

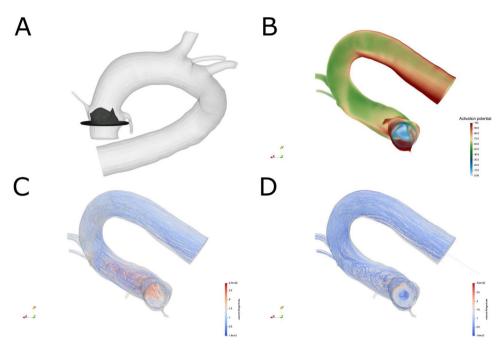


Figure 1. A: three-dimensional representation of the valve in its implantation plane based on computed tomography of one of the patients and the valve designed by the authors. B: platelet activation potential calculated at peak systole for the linear inflow profile. C: current lines in the transvalvular flow model for the linear (uniform) velocity profile in the left ventricular outflow tract. D: streamlines in the transvalvular flow model with helical flow. WSS, wall shear stress.

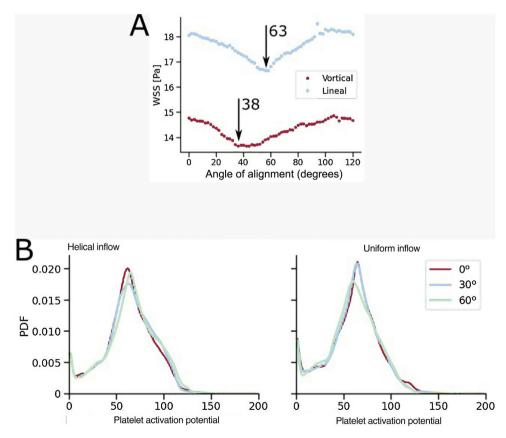


Figure 2. A: fluid dynamic simulation of energy efficiency of blood outflow through the valve (the lower the value, the higher the efficiency) as a function of the degree of alignment (shown in degrees) and the type of simulation (uniform or helical). The upper dashed line (blue) shows higher efficiency in severe misalignment (63°) if the flow is linear; the lower dashed line (red) shows higher overall efficiency and, in particular, in the absence of moderate or severe misalignment (38°) if the flow is helical. B: simulation of the degree of platelet activation as a function of the degree of commissural alignment (0°: perfect alignment; 30°: mild misalignment; 60°: severe misalignment), with no difference between helical (left) or linear (right) flow. PDF, probability density function.

Progressive understanding of the relevance of commissural alignment in TAVI has led to the development of custom software to help plan this procedure and to technical changes in the devices that make the procedure easier and faster. On the other hand, it is known that platelet activation takes place not only in situations of increased wall stress, such as aortic stenosis, but also in diseases with abnormal flows, such as bicuspid aortic valve disease.⁵ An association has been found between platelet activation and an increased risk of thrombotic and hemorrhagic phenomena.⁵ Previous studies have shown that computational models are quite accurate in predicting such activation; for example, they are able to predict thrombosis in coronary bifurcation stenting (figure 1B).³ The absence of differences in the degree of platelet activation as a function of the degree of misalignment, regardless of whether the flow is helical (figure 2B, left) or linear (figure 2B, right), suggests that the risk of clinical or subclinical thrombosis of the leaflets does not appear to differ as a function of the degree of misalignment. These models are based on purely mechanical assumptions (blood velocity fields), although there is a possibility that the development of more advanced models, which include elements of the coagulation cascade, will enable the detection of differences in thrombogenicity depending on commissural alignment.

On the other hand, the impact on ventricular energy efficiency suggests that, in each individual patient, the result in terms of gradients could differ depending on the degree of alignment. To detect this difference, the models used should simulate flows that are as realistic as possible (helical), given that current simulation laboratories use uniform flows through the prostheses under study.¹ Should our findings be prospectively validated, it could be demonstrated that commissural alignment has an impact on prosthetic valve durability.

In conclusion, CT analysis prior to TAVI is highly accurate in predicting the rotation of the system to achieve correct commissural alignment, especially when using self-expanding prostheses. Furthermore, based on the same imaging tests, computational fluid dynamics analysis highlights the benefits of commissural alignment in taking advantage of the mechanical energy provided by the ventricular wall, but which can only be visualized in realistic (helical) simulations of flow in the outflow tract. We found that commissural alignment had no impact on the risk of leaflet thrombosis.

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Potential for controlled heart donation after circulatory determination of death

Potencialidad de la donación cardiaca de donantes en asistolia controlada

To the Editor,

Heart transplant is the treatment of choice for advanced heart failure. The number of ideal brain-dead donors, however, has

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AUTHORS' CONTRIBUTIONS

I. Amat-Santos and J. Sierra-Pallares designed and conducted the study. All authors approved the final version of the manuscript.

CONFLICTS OF INTEREST

None declared.

APPENDIX. SUPPLEMENTARY DATA

Supplementary data associated with this article can be found in the online version, at https://doi.org/10.1016/j.rec.2023.01.012

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REFERENCES

- 1. Hatoum H, Dollery J, Lilly SM, Crestanello JA, Dasi LP. Implantation depth and rotational orientation effect on valve-in-valve hemodynamics and sinus flow. *Ann Thorac Surg.* 2018;106:70–78.
- Khan JM, Rogers T, Weissman G, et al. Anatomical characteristics associated with hypoattenuated leaflet thickening in low-risk patients undergoing transcatheter aortic valve replacement. *Cardiovasc Revasc Med.* 2021;27:1–6.
- García-Carrascal P, García-García J, Sierra-Pallares J, et al. Numerical study of blood clots influence on the flow pattern and platelet activation on a Stented Bifurcation Model. Ann Biomed Eng. 2017;45:1279–1291.
- Redondo A, Valencia-Serrano F, Santos-Martínez S, et al. Accurate commissural alignment during ACURATE neo TAVI procedure. Proof of concept. *Rev Esp Cardiol.* 2022;75:203–212.
- Raschpichler M, Flint N, Yoon SH, et al. Commissural Alignment After Balloon-Expandable Transcatheter Aortic Valve Replacement Is Associated With Improved Hemodynamic Outcomes. *JACC Cardiovasc Interv*. 2022;15:1126–1136.

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decreased in recent decades, requiring the implementation of strategies to increase the number of transplantable hearts. These strategies include expanding donor acceptability criteria and using controlled donation after circulatory determination of death (cDCD), which currently accounts for 40% of organ donations in Spain.¹

Promising results have been reported for different cDCD protocols implemented in Spain since 2020. No studies to date, however, have analyzed the potential for cDCD in Spain. To this end, we conducted a retrospective, observational study of all