



# **Virtual cohort generation of Aortic Valve Stenosis geometries**

## **Sabine Verstraeten**

Frans van de Vosse

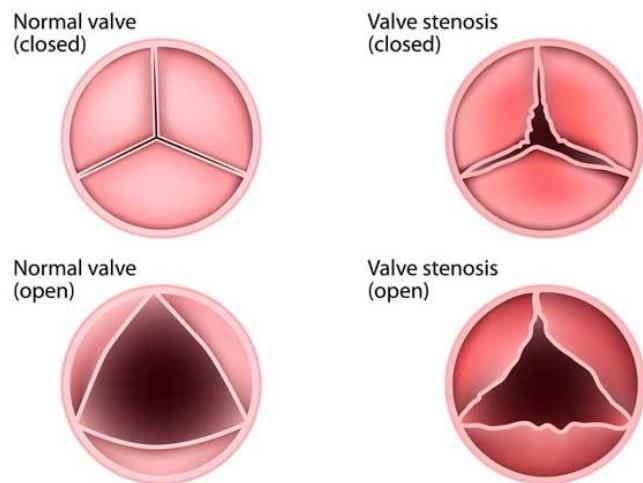
Wouter Huberts

ESB 09/07/2023

[s.c.f.p.m.verstraeten@tue.nl](mailto:s.c.f.p.m.verstraeten@tue.nl)

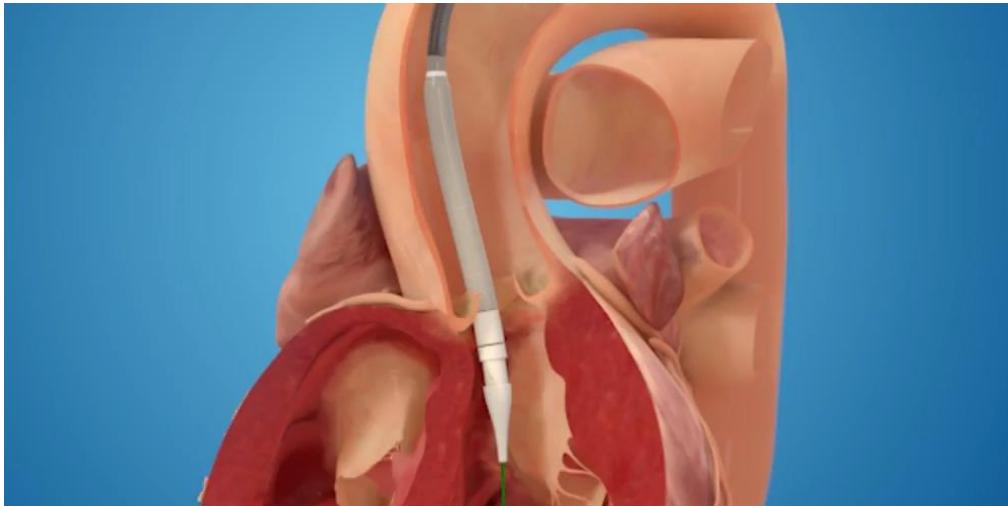


# Aortic valve stenosis



Heart-valve-surgery.com (2021)

# Transcatheter aortic valve implantation (TAVI)



## Benefits

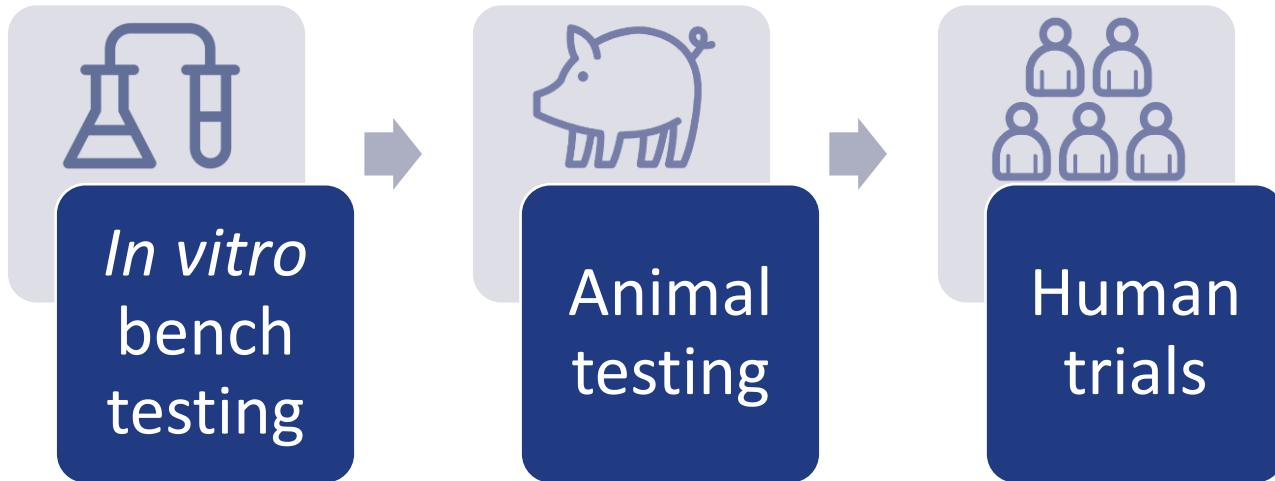
- Relief from symptoms
- Increased life expectancy
- Accessible for all patients



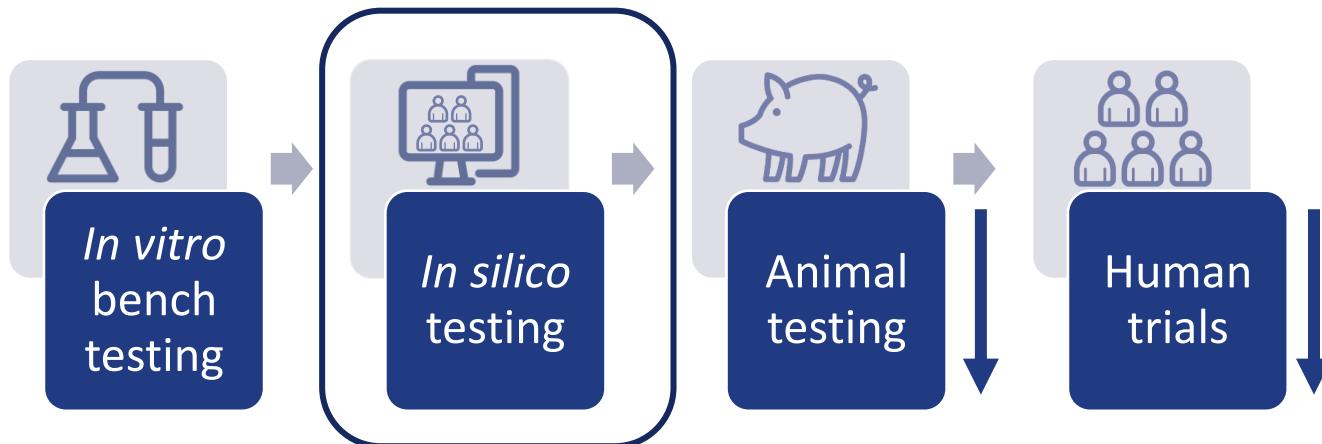
## Risks

- Paravalvular leakage
- Thrombosis
- Conduction problems

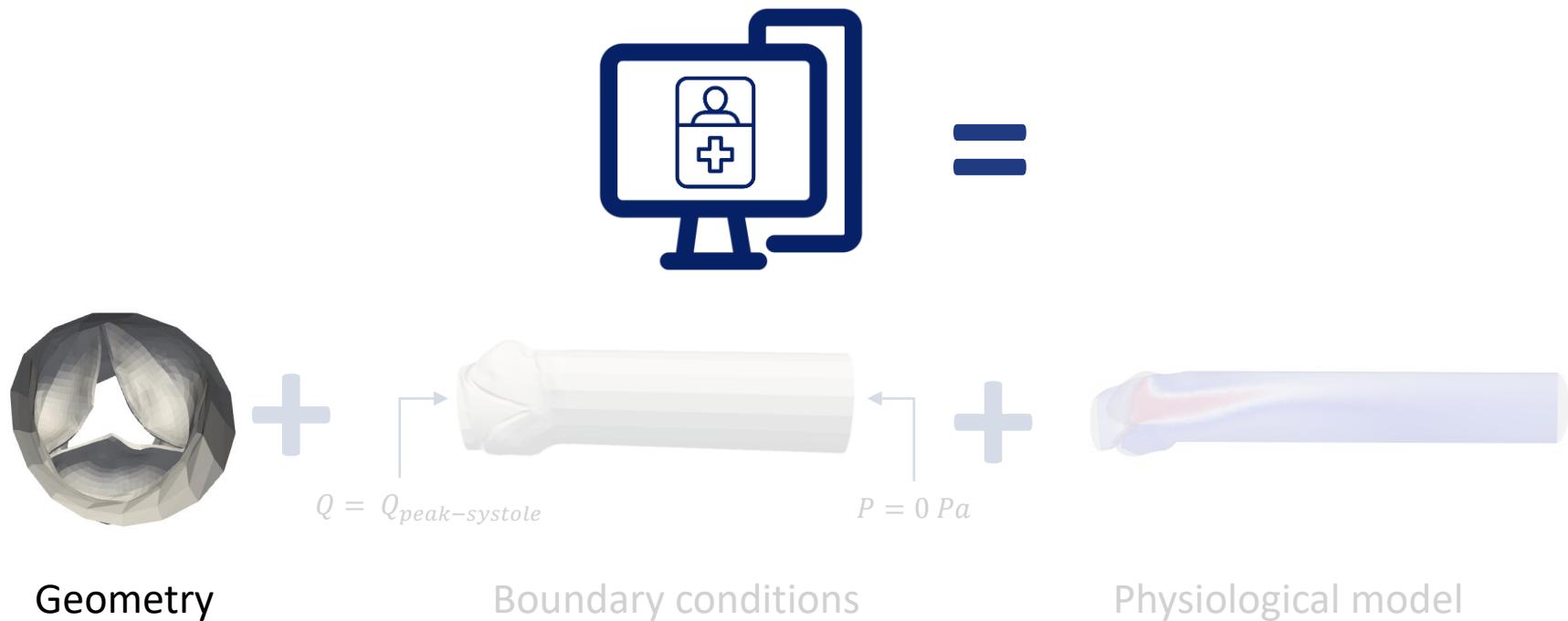
# Developing & testing TAVI devices



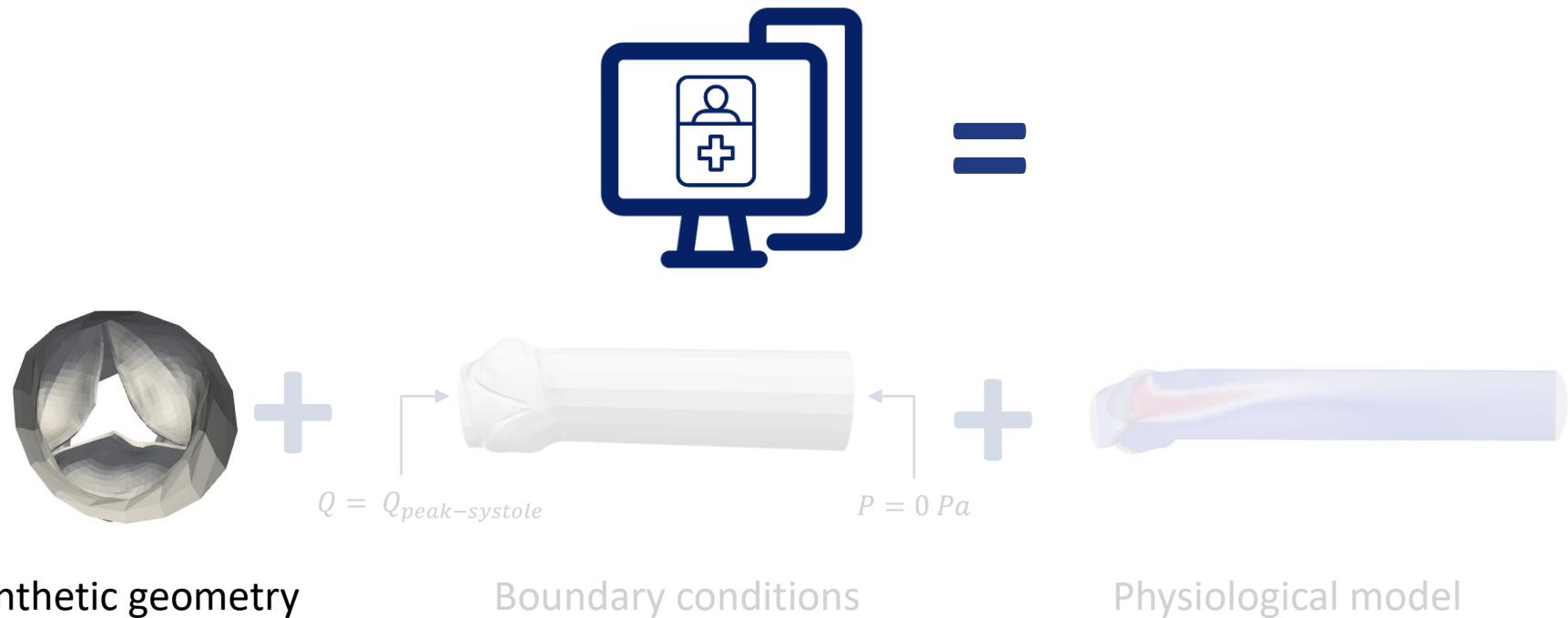
# Developing & testing TAVI devices



# The virtual patient

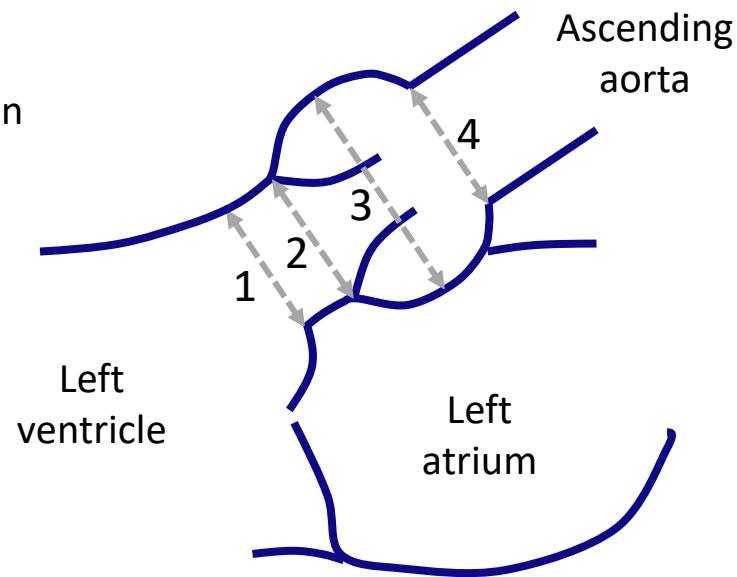


# The virtual patient



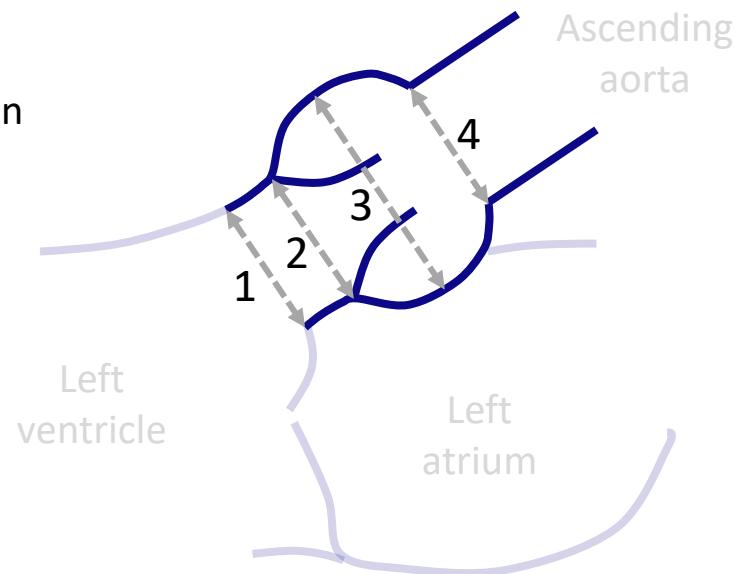
# Aortic valve geometry

1. LVOT
2. Annulus
3. Sinuses of Valsalva
4. Sinotubular junction



# Aortic valve geometry

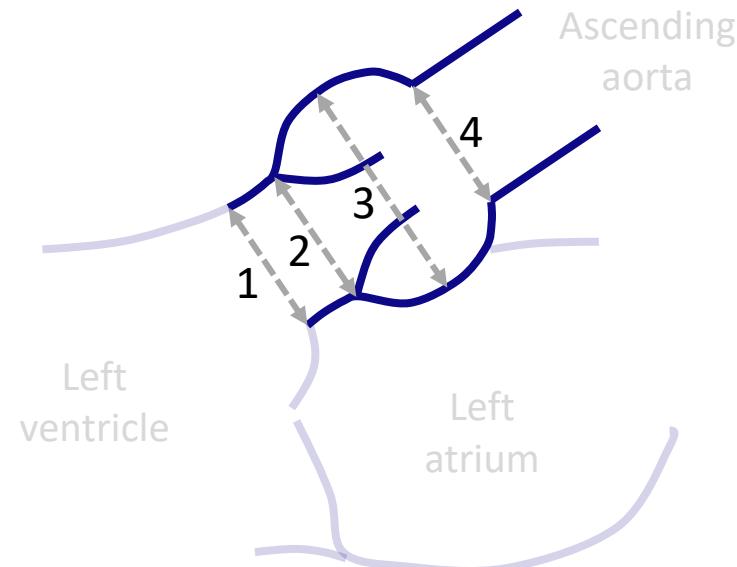
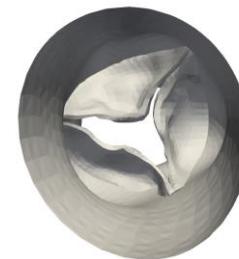
1. LVOT
2. Annulus
3. Sinuses of Valsalva
4. Sinotubular junction



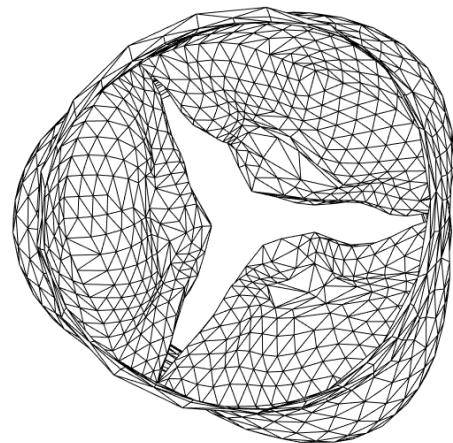
# Aortic valve geometry

Real patient data set

- 97 geometries
- Peak-systolic
- Severe stenosis



# Statistical shape modelling (SSM)

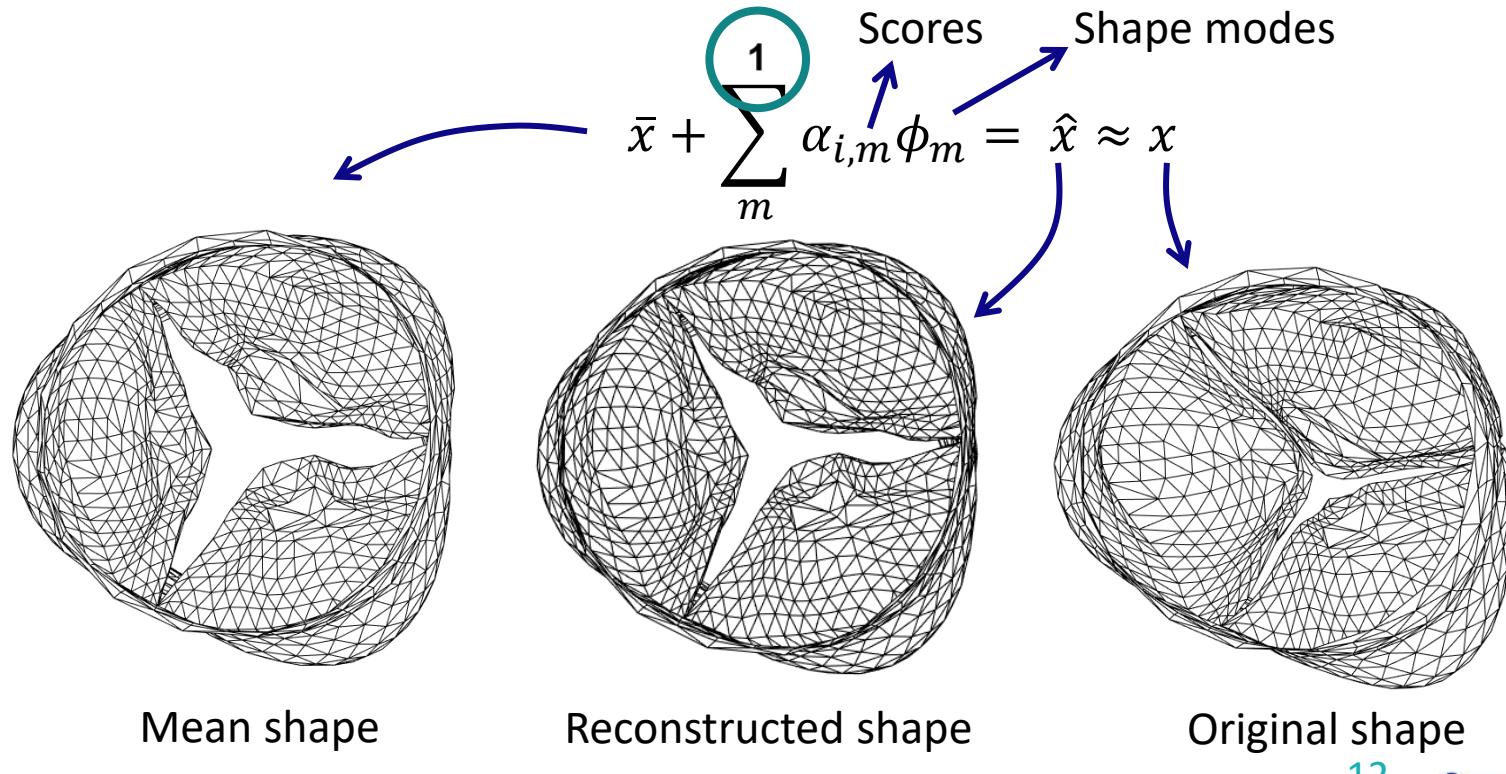


Mean shape

$$\bar{x} + \sum_m^N \alpha_m \phi_m = \hat{x} \approx x$$

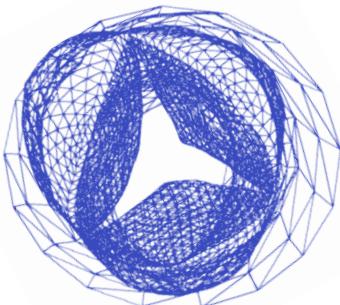
A mathematical equation illustrating the decomposition of a shape. On the left, a curved arrow points from the mean shape to the equation. The equation itself is  $\bar{x} + \sum_m^N \alpha_m \phi_m = \hat{x} \approx x$ . A blue arrow points from the term  $\alpha_m$  to the label "Scores". Another blue arrow points from the term  $\phi_m$  to the label "Shape modes".

# Statistical shape modelling (SSM)

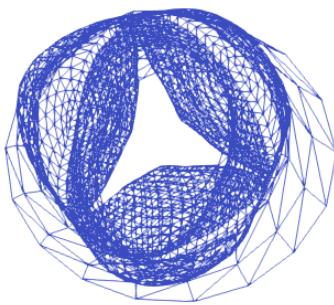


# Statistical shape modelling (SSM)

Conventional SSM

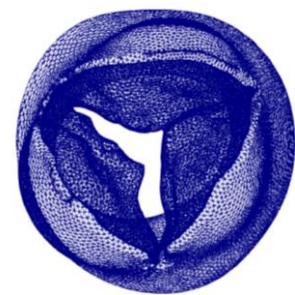


=

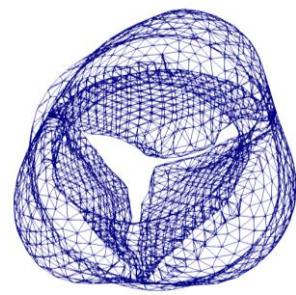


Inter-patient topology required!

Non-parametric SSM



≠



24254 points

4063 points



Applicable to all sets of geometries!

# Synthetic geometries with non-parametric SSM

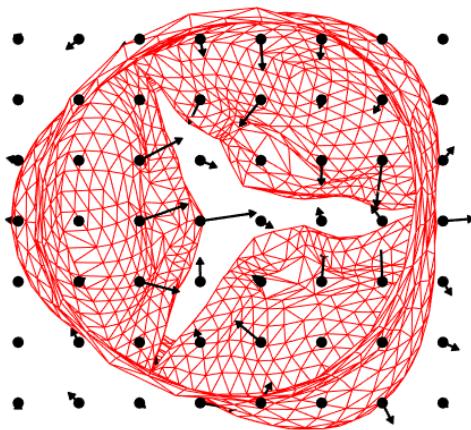


# Synthetic geometries with non-parametric SSM



# Non-parametric representation

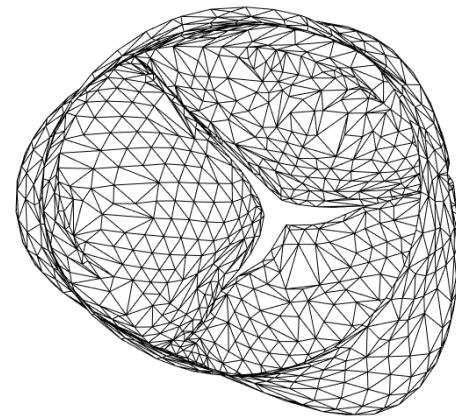
Template  $\bar{x}$



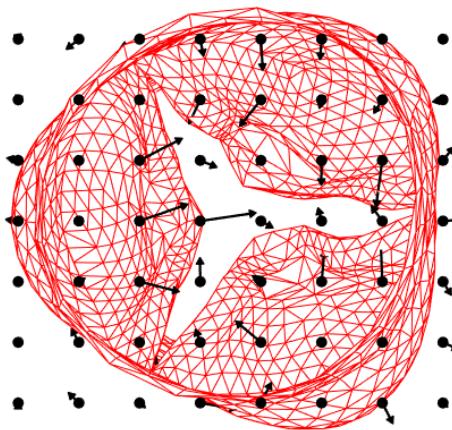
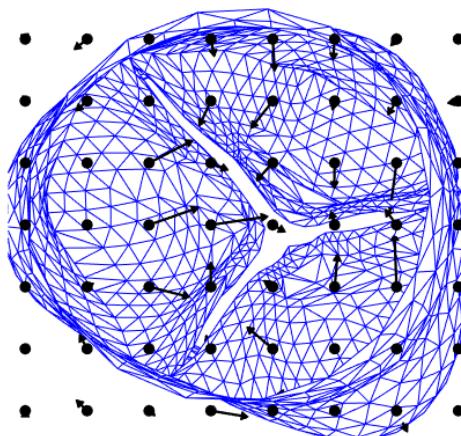
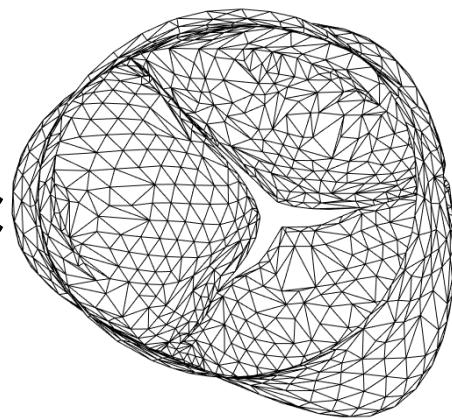
$N$  deformation vectors  $\beta_k$

$$\bar{x} + \sum_k^N K(\bar{x}, c_k) \cdot \beta_k$$

Patient  $x$



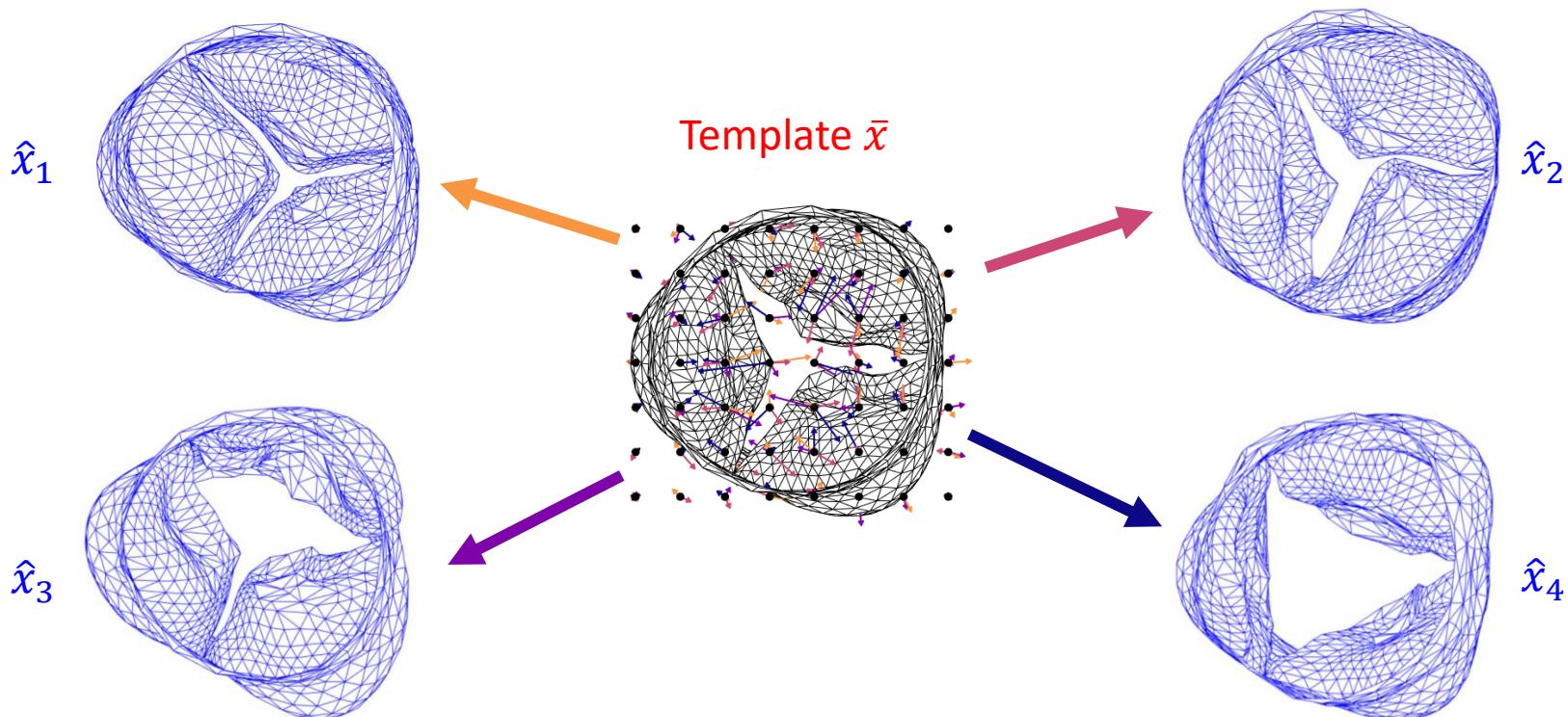
# Non-parametric representation

Template  $\bar{x}$ Deformed template  $\hat{x}$ Patient  $x$ 

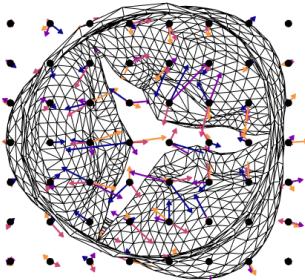
$N$  deformation vectors  $\beta_k$

$$\bar{x} + \sum_k^N K(\bar{x}, c_k) \cdot \beta_k = \hat{x} \approx x$$

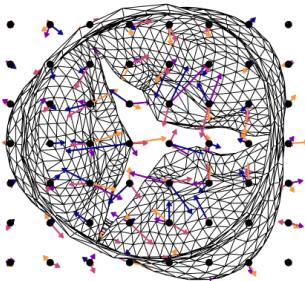
# Non-parametric representation



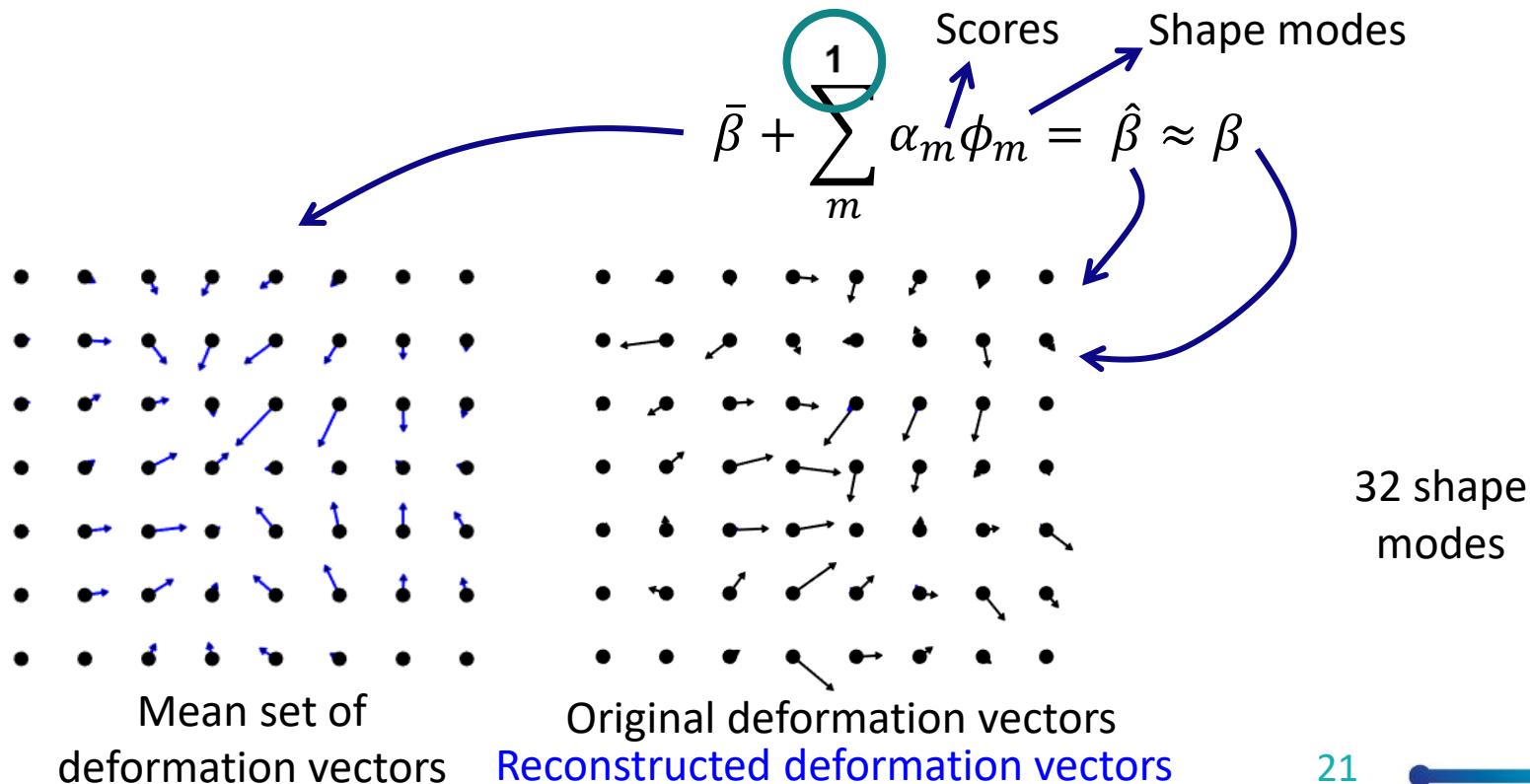
# Synthetic geometries with non-parametric SSM



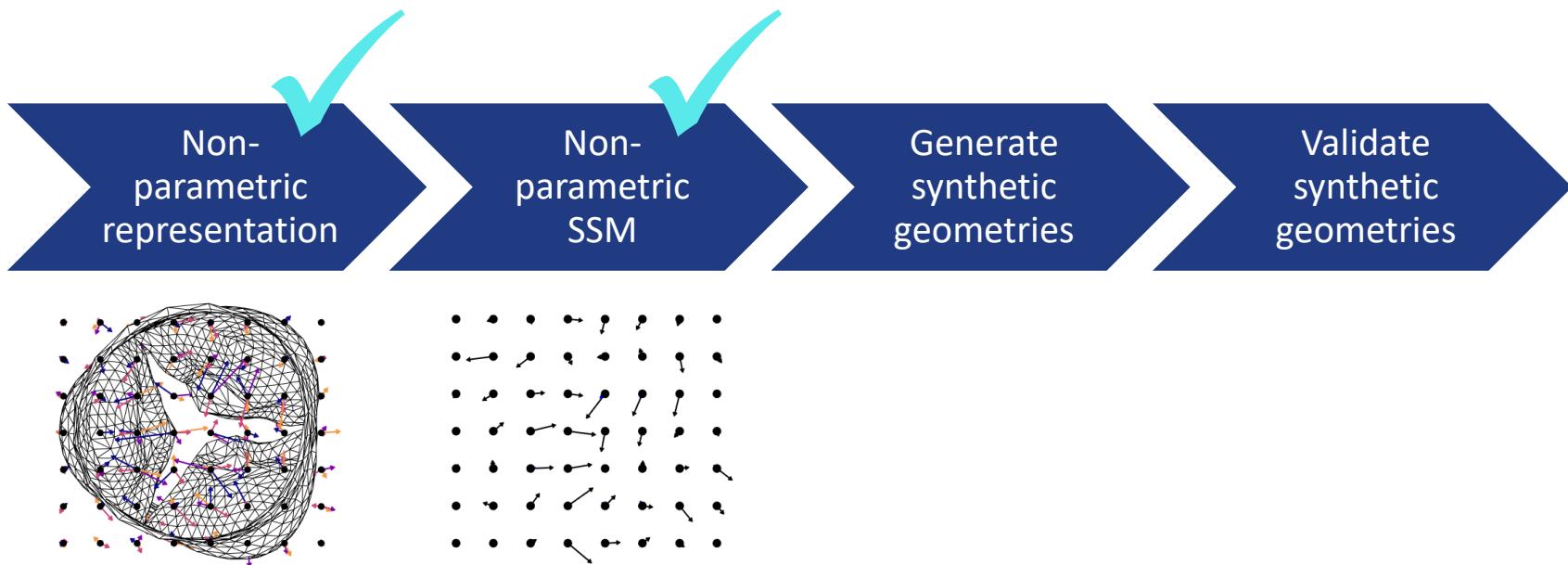
# Synthetic geometries with non-parametric SSM



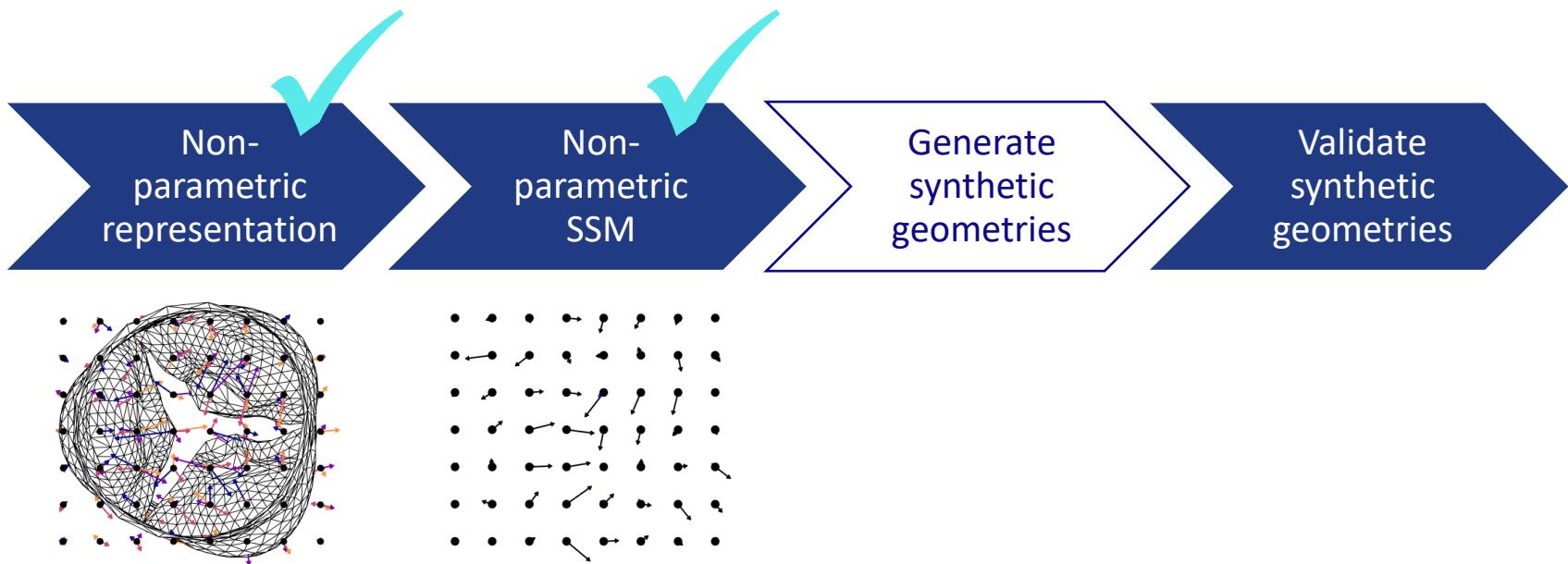
# Non-parametric SSM



# Synthetic geometries with non-parametric SSM



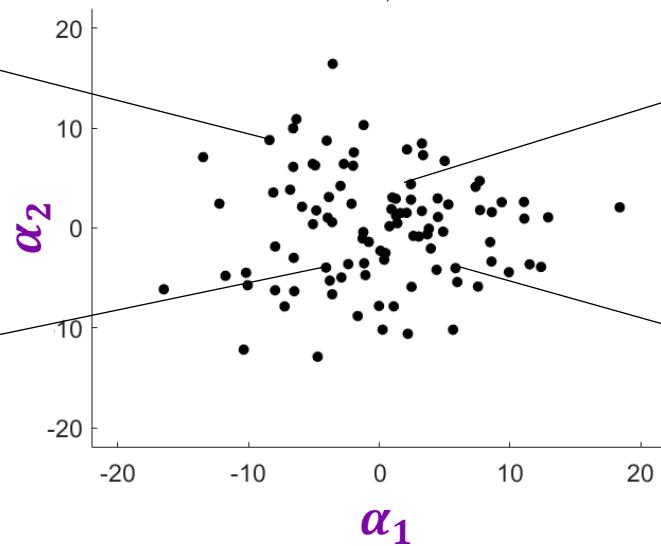
# Synthetic geometries with non-parametric SSM



# Generation of synthetic geometries

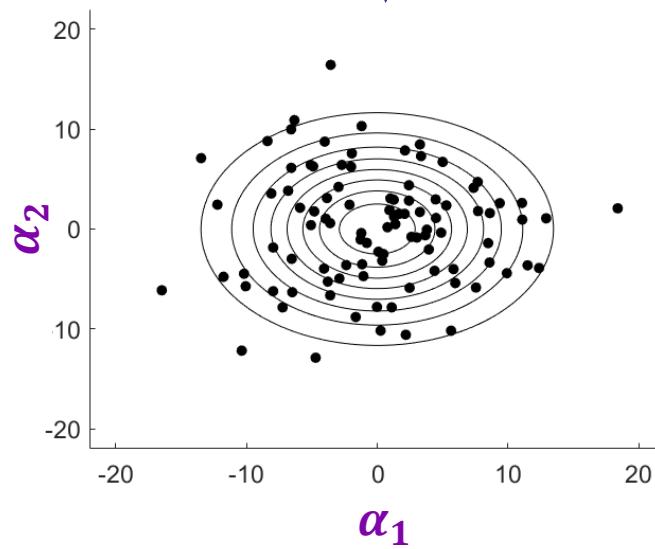


$$\bar{\beta} + \sum_m^{32} \alpha_m \phi_m = \hat{\beta}$$

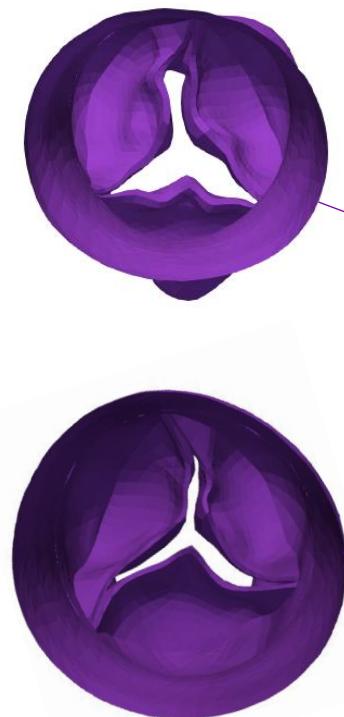


# Generation of synthetic geometries

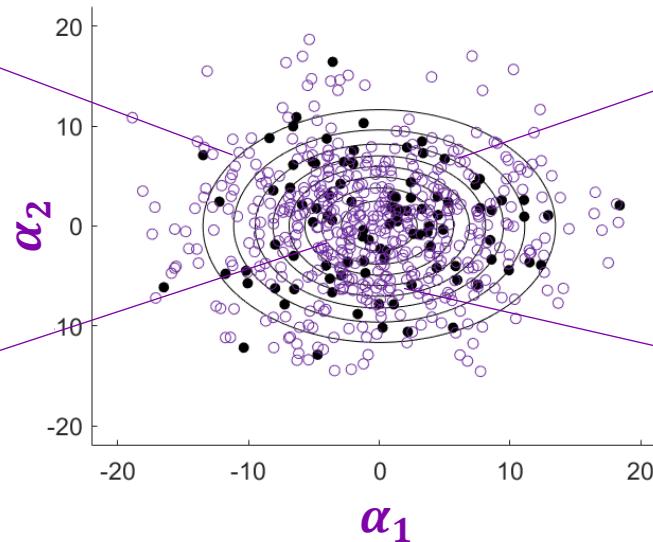
$$\bar{\beta} + \sum_m^{32} \alpha_m \phi_m = \hat{\beta}$$



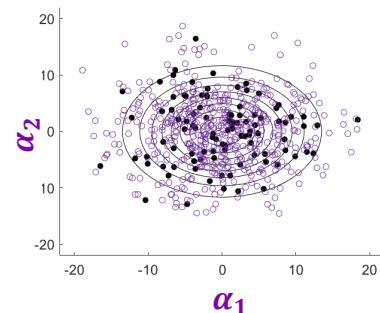
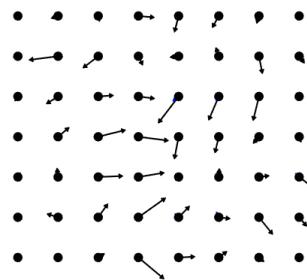
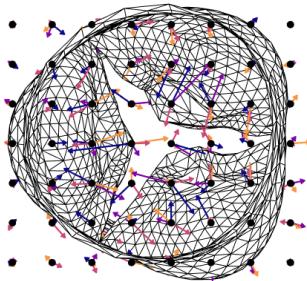
# Generation of synthetic geometries



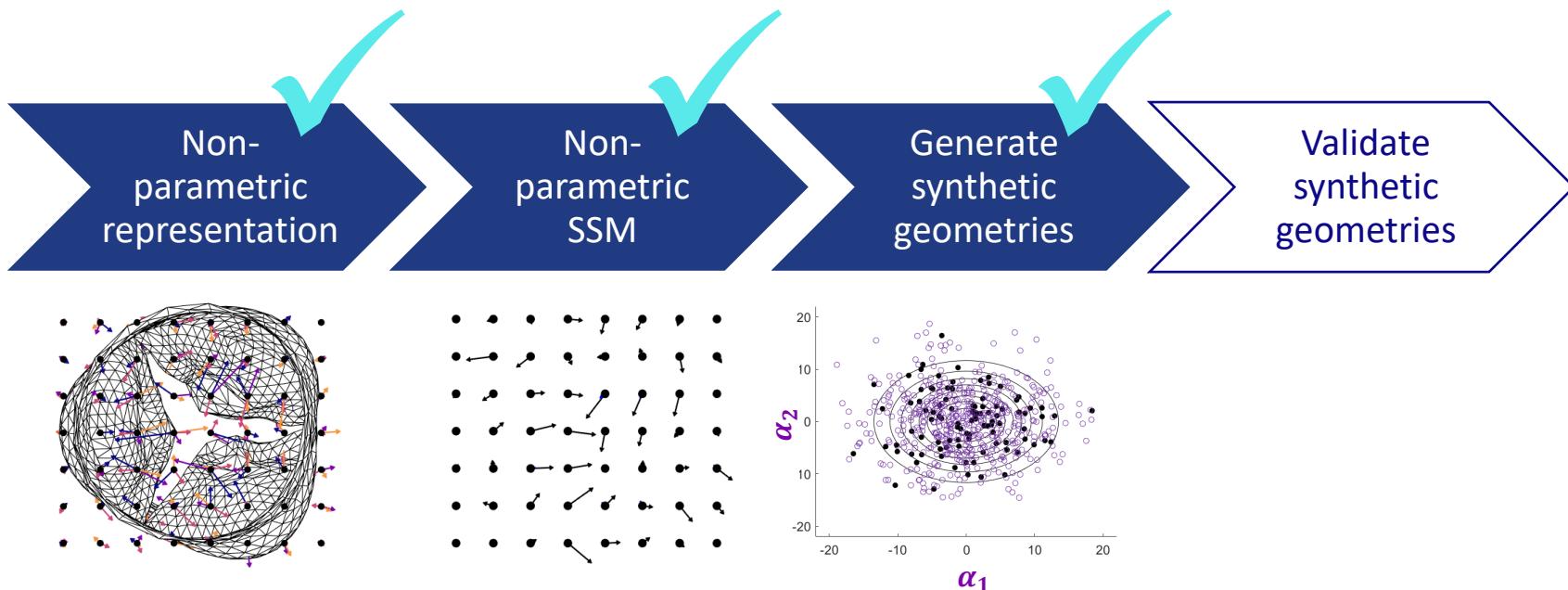
$$\bar{\beta} + \sum_m^{32} \alpha_m \phi_m = \hat{\beta}$$



# Synthetic geometries with non-parametric SSM

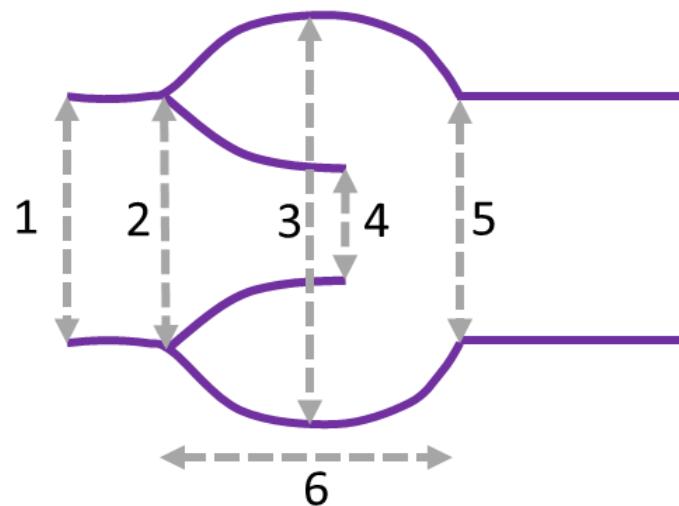


# Synthetic geometries with non-parametric SSM



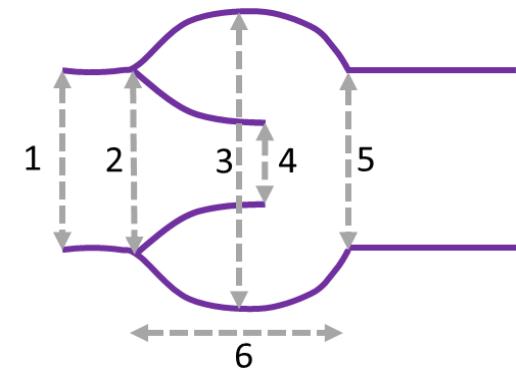
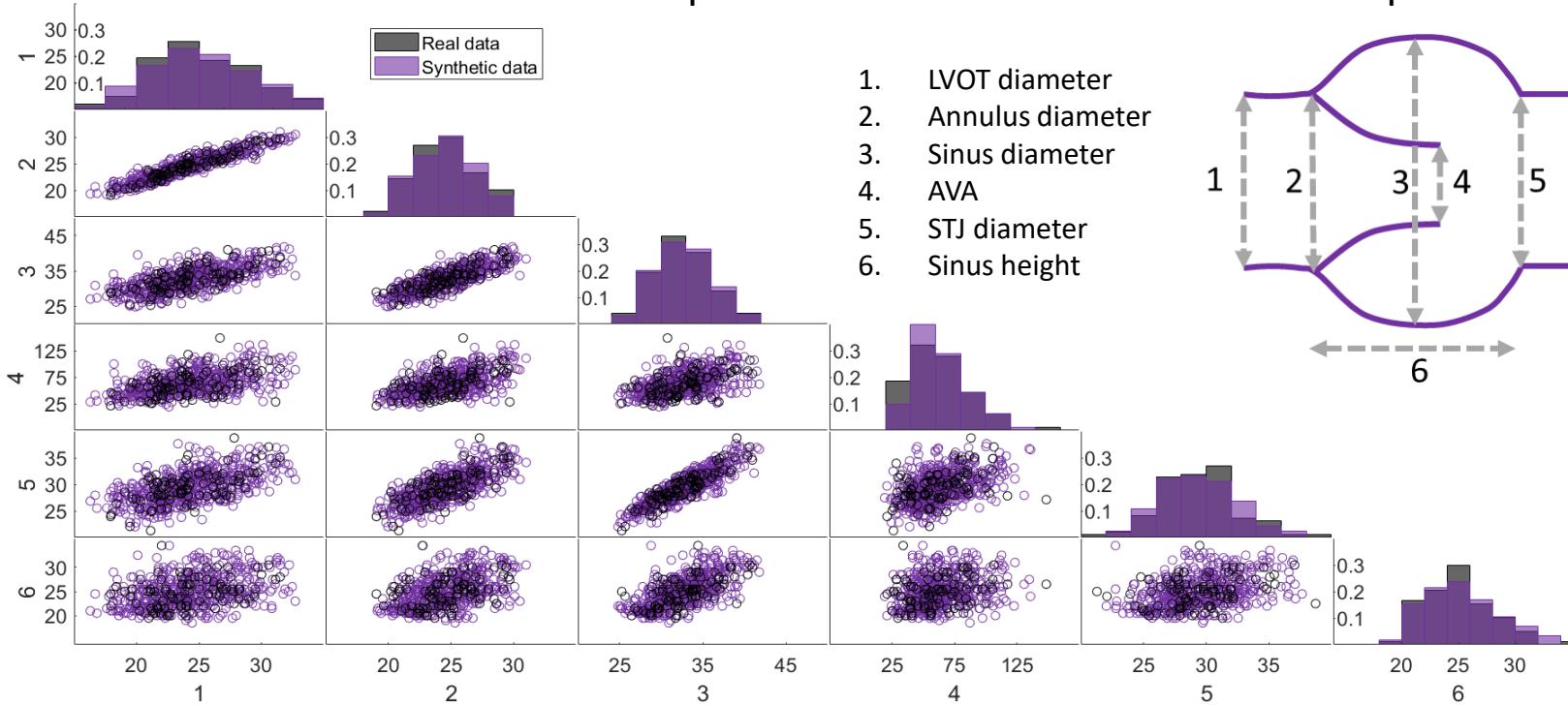
# Validation

1. LVOT diameter
2. Annulus diameter
3. Sinus diameter
4. AVA
5. STJ diameter
6. Sinus height



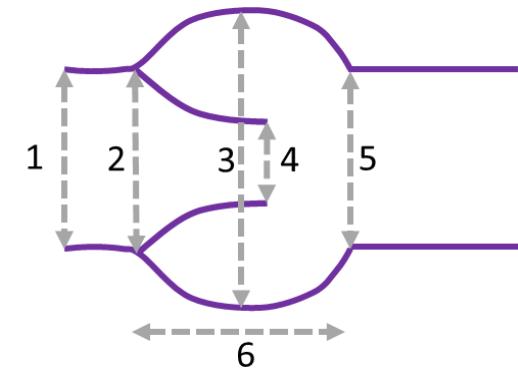
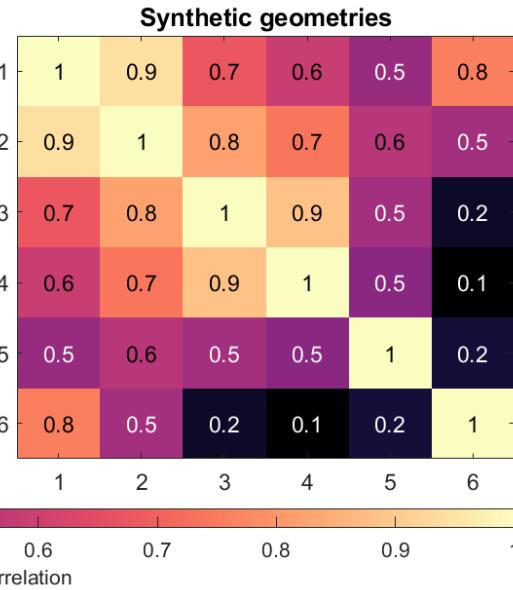
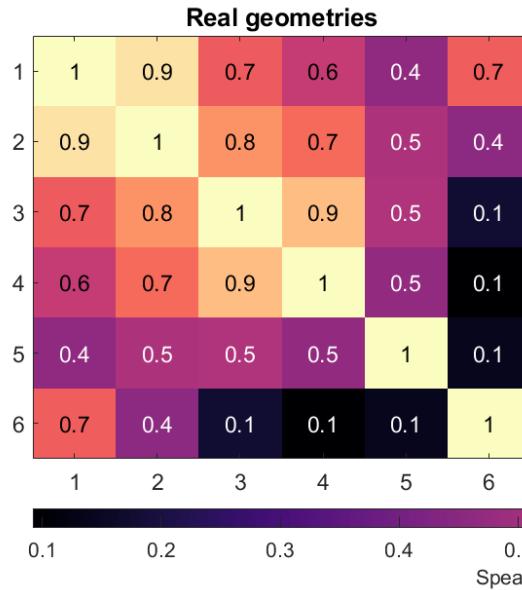
# Validation

Non-parametric multivariate ANOVA test:  $p = 0.86 > 0.05$



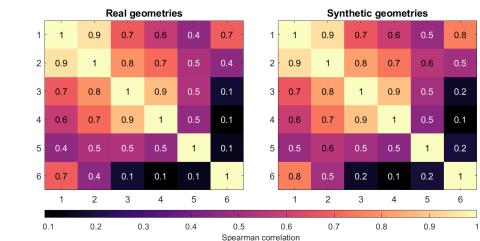
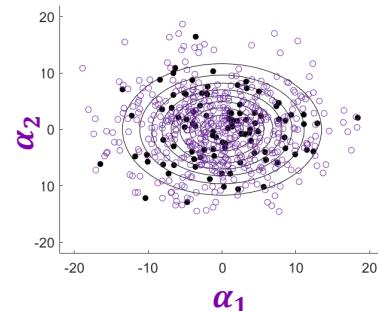
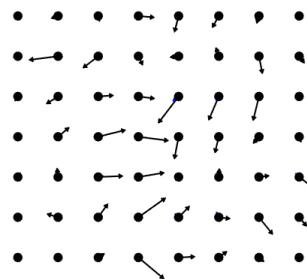
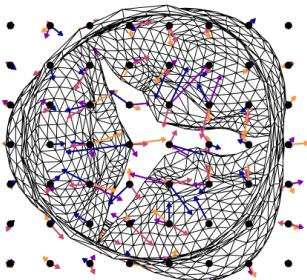
# Validation

Non-parametric multivariate ANOVA test:  $p = 0.86 > 0.05$



1. LVOT diameter
2. Annulus diameter
3. Sinus diameter
4. AVA
5. STJ diameter
6. Sinus height

# Synthetic geometries with non-parametric SSM



# Virtual cohort generator



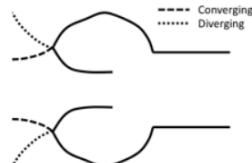
**SIMCor**

Virtual cohort generator to generate synthetic stenosed aortic valve geometries

Enter desired number of virtual cases

Choose template type

Choose desired LVOT shape



Choose range of angles ( $a$ ) between LVOT and ascending aorta

Set angle range (between 0 and 70 degrees)

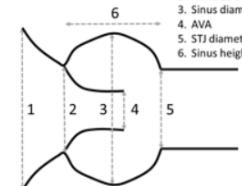
Enter minimum

Enter maximum

Extract anatomical features

Anatomical features

1. LVOT diameter
2. Annulus diameter
3. Sinus diameter
4. AVA
5. STJ diameter
6. Sinus height



Enter desired number of workers in parallel pool (default 1)

Start

# Virtual cohort generator

**SIMCor** Virtual cohort generation for *in silico* trials of transcatheter aortic valve implantation (TAVI)

“S.C.P.M. Verstraeten\*, N.J.M.M. Hoedjik\*, P.A.L. Tonino, J. Brink, M. Capelli, F.N. de Vosse, M.W. Huberts, on behalf of SIMCor consortium”  
Eindhoven University of Technology, Maastricht University, Erasmus University Rotterdam, University College London, Radboud University

**Need for *in silico* trials**  
*In silico* trials can speed up the long and expensive process of developing cardiovascular implantable devices, by integrating them in the validation chain:

- Designing
- In vitro tests
- Animal tests
- Clinical trials

During *in silico* trials devices are tested on cohorts of virtual patients. Virtual patients are computer models that simulate human physiology and response after device implantation.

**In Silico trials to investigate TAVI complications**  
This study aims to generate synthetic aortic valve stenosis geometries related to the shape of the left ventricular outflow tract (LVOT) [1] and the angle between the LVOT and the ascending aorta [2] (see next figure).

**Example applications of virtual cohort generator**

Industry, Academy, Clinic

LVOT-shape filter: Logistic regression model that predicts chance of valve stenosis, based on shape coefficients.

Sampling: Geometries are generated by 3D shape sampling, resulting from a nonparametric model [2], based on 57 real patient geometries. Synthetic geometries are sampled from a 3D distribution fitted to real data.

Angle filter: Linear regression model that predicts if the angle is in the normal range, based on angle coefficients.

Validation: 1. LVOT diameter  
2. Annulus diameter  
3. Sinus diameter  
4. AVA  
5. Aortic diameter  
6. Sinus height

Computational fluid dynamics (CFD) and fluid-structure interaction (FSI) models can calculate valve dynamics before and after device implantation, using anatomical features and synthetic data as source and reconstructing geometries from imaging data is time-consuming. Using synthetic geometries is a promising solution.

Aim:  
1. Aim of this study is to develop a virtual cohort generator that is:  
a. Able to generate physiologically plausible, synthetic aortic valve stenosis geometries.  
b. Allows for the selection of clinically relevant anatomical features.

Scan for poster video

Poster session III: 12/07 13.15 – 14.15



Virtual cohort generator to generate synthetic stenosed aortic valve geometries

Enter desired number of virtual cases:

Choose template type: Double layer leaflets

Choose desired LVOT shape: No preference

Choose range of angles ( $\alpha$ ) between LVOT and ascending aorta  
Set angle range (between 0 and 70 degrees)  
Enter minimum:   
Enter maximum:

Extract anatomical features

Anatomical features:  
1. LVOT diameter  
2. Annulus diameter  
3. Sinus diameter  
4. AVA  
5. Aortic diameter  
6. Sinus height

Enter desired number of workers in parallel pool (default 1):

**Start**

Submitted paper: generation of synthetic aortic valve stenosis geometries for *in silico* trials  
(Verstraeten et al. 2023)

Special thanks to

Charité  
Jan Brüning

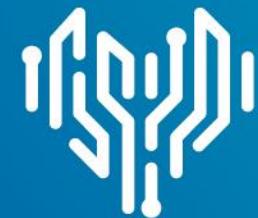


University college of London  
Claudio Capelli, Jan Bruse



And all other SIMCor partners

Ansys  
Martijn Hoeijmakers



Thanks!

Sabine Verstraeten  
[s.c.f.p.m.verstraeten@tue.nl](mailto:s.c.f.p.m.verstraeten@tue.nl)

Scan for poster video



This project has received funding from the European Union's Horizon 2020 research and innovation programme under grant agreement N° 101017578