



## Generative models

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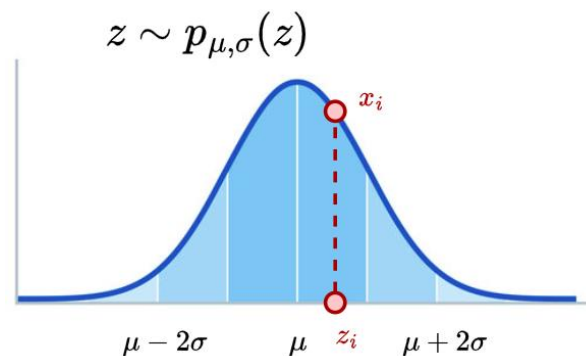
# What is the interest of generative models ?

## ► How to generate synthetic faces ?



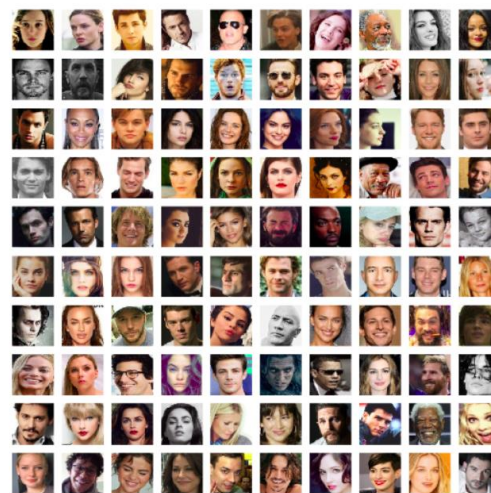
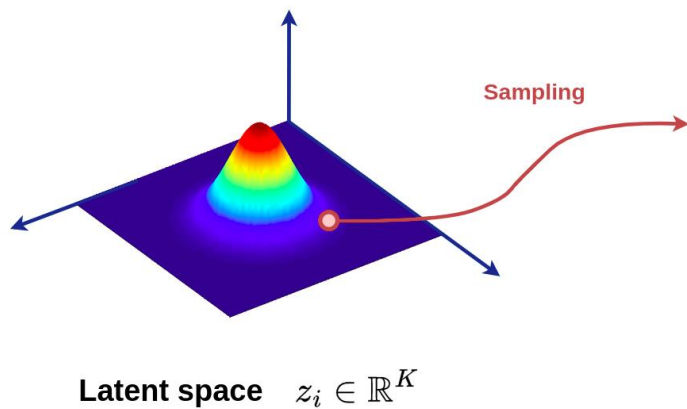
By modeling the corresponding distribution  $p_{\theta}(\cdot)$  !

► Reminder: normal distribution



# What are the interest of generative models ?

## ► How to model complex distributions ?

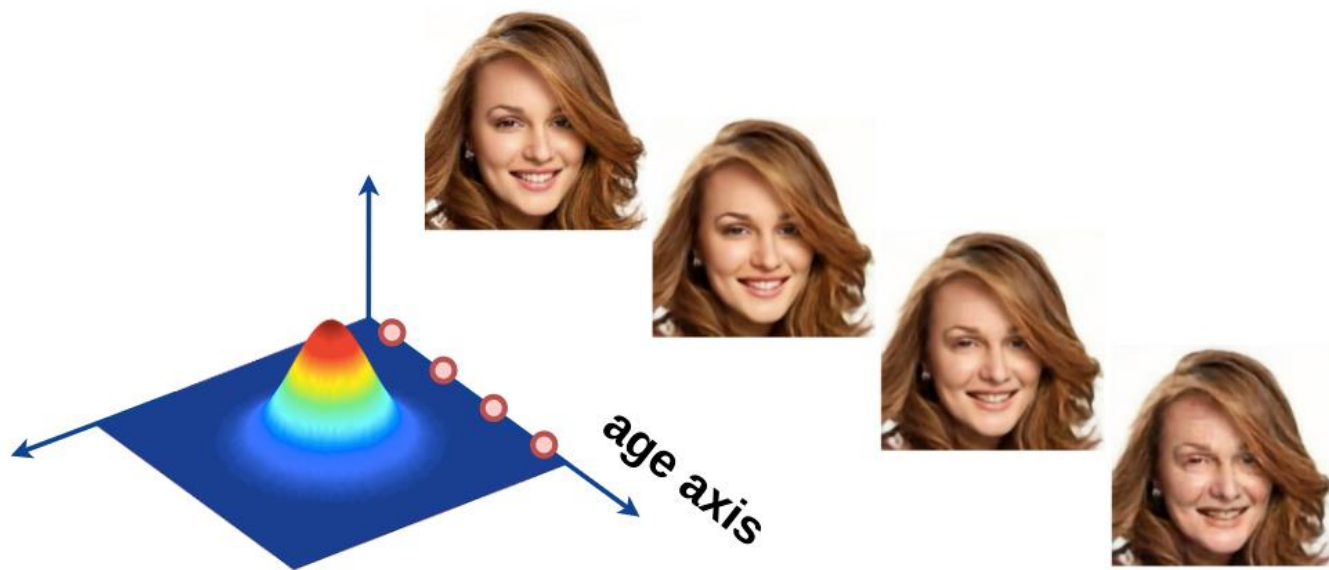


Face distribution

# What are the interest of generative models ?

► What for ?

One obsession is to master the latent space !!!

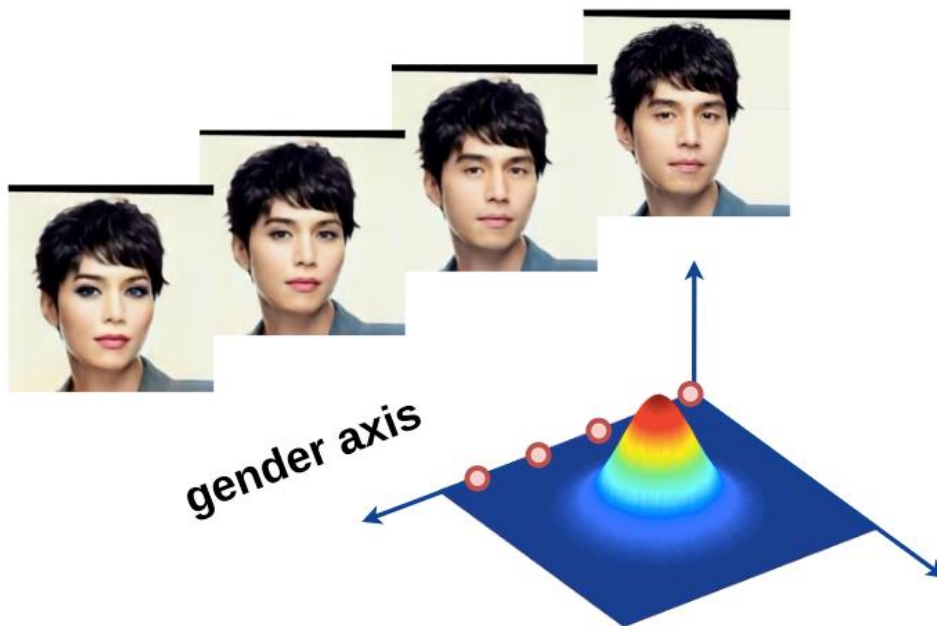


Latent space  $z_i \in \mathbb{R}^K$

# What are the interest of generative models ?

► What for ?

One obsession is to master the latent space !!!



Latent space  $z_i \in \mathbb{R}^K$

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# Auto-encoders

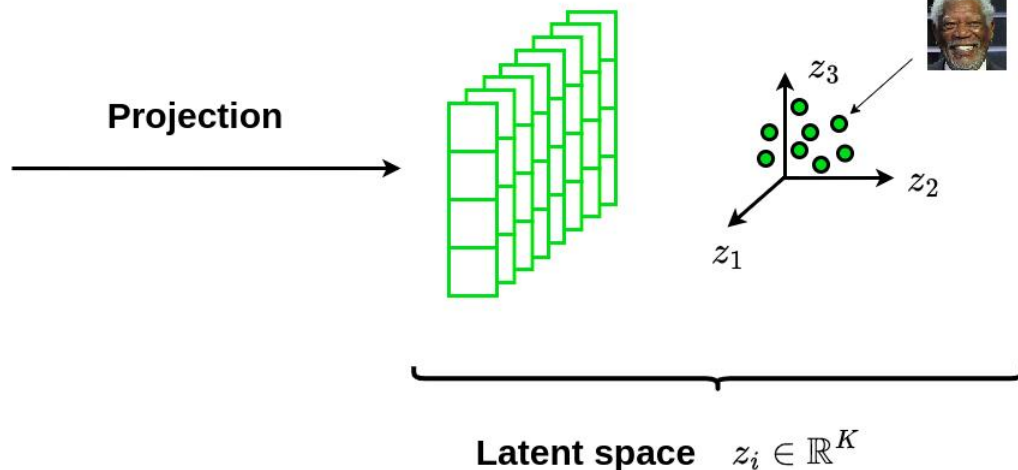
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# How to learn a distribution ?

## ► Projection into a simpler, lower-dimensional representation space

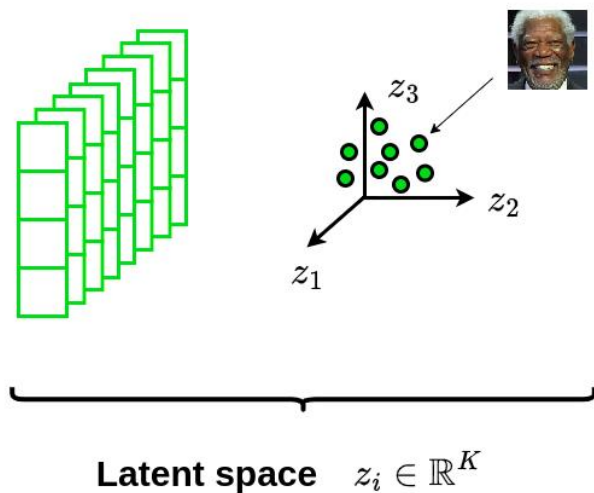


Input space  $x_i \in \mathbb{R}^{N \times M}$



# How to learn a complex distribution ?

## ► How to have a relevant representation space ?



Reconstruction  
→

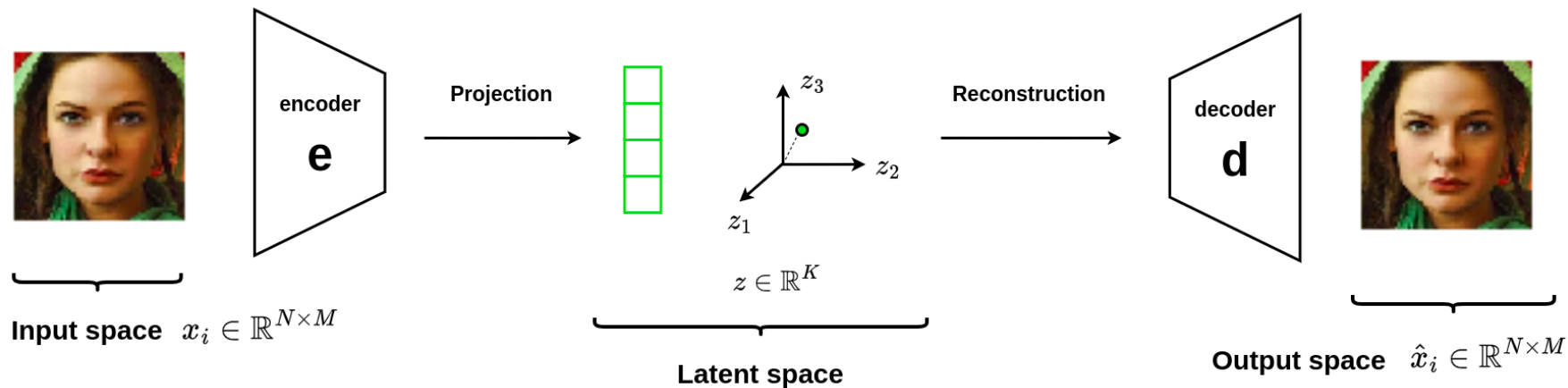


Output space  $\hat{x}_i \in \mathbb{R}^{N \times M}$



# Auto-encoder framework

## ► Standard architecture

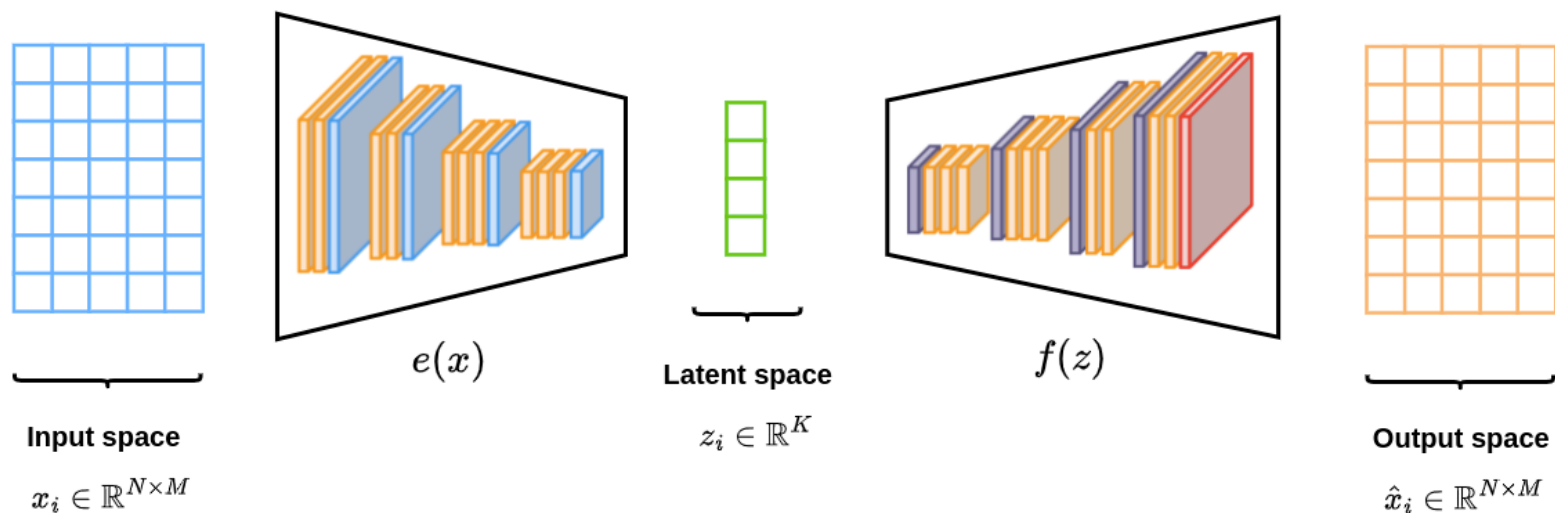


## ► Deep learning loss function

$$\text{loss} = \|x - \hat{x}\|^2$$

# Deep learning implementation

## ► Encoder / Decoder modeled through (convolutional) neural networks

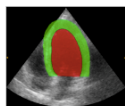


$$\text{loss} = \|x - f(e(x))\|^2$$

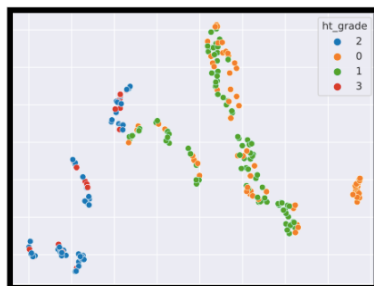
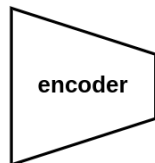
# Interest of auto-encoders

## ▶ Auto-encoder ? For what purpose ?

### → Data representation

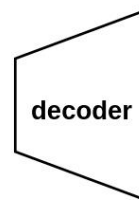


Patients



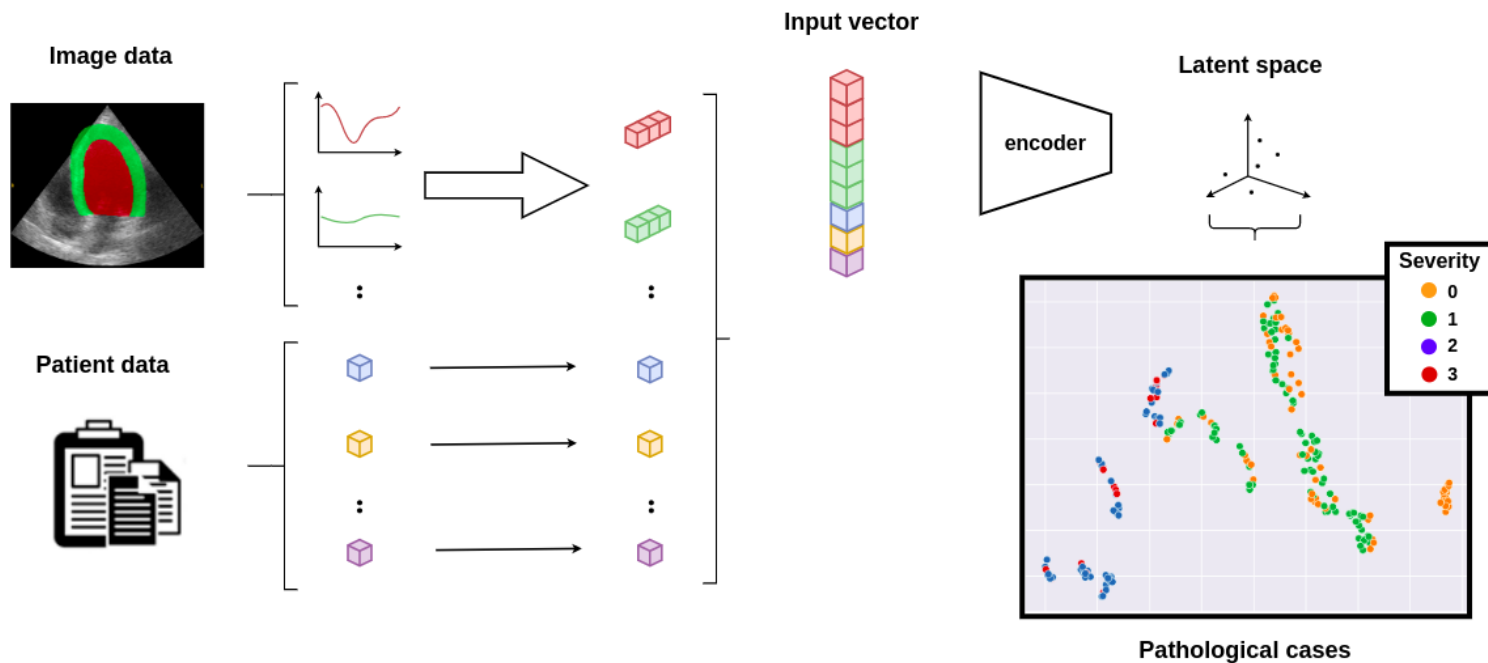
Population representation

### → Generative model



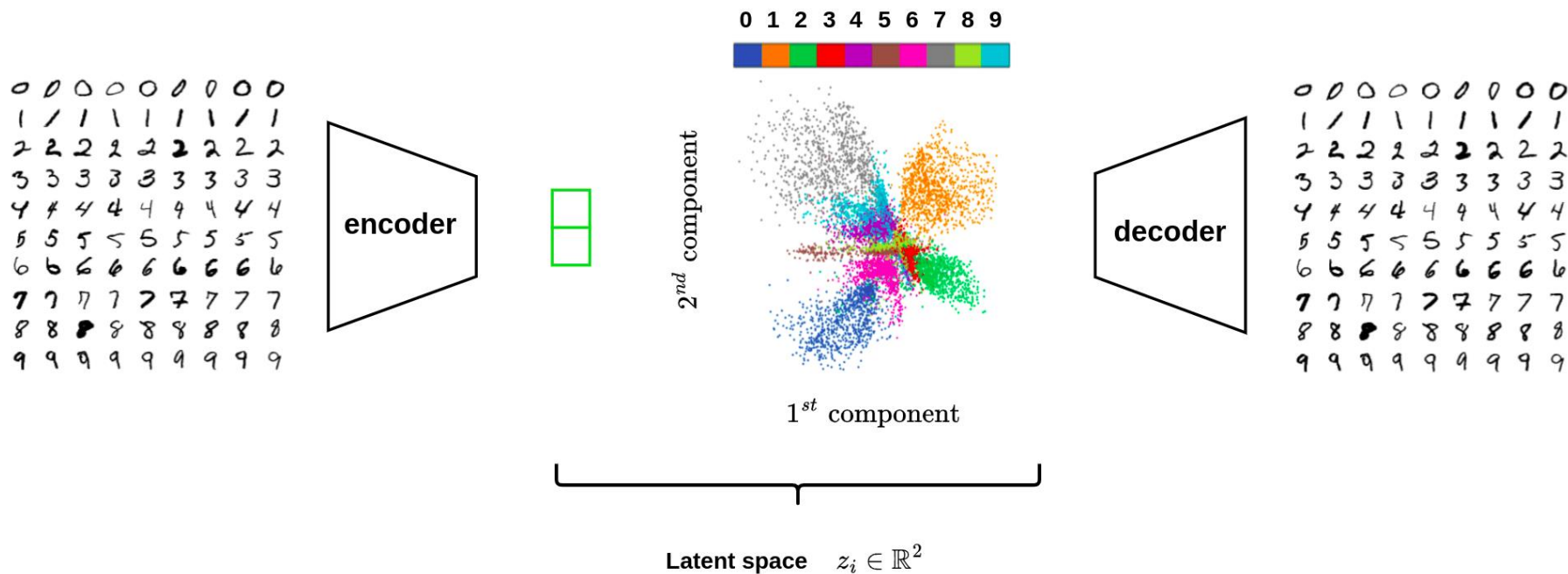
0	0	0	0	0	0	0	0	0
1	/	/	\	/	/	\	/	/
2	2	2	2	2	2	2	2	2
3	3	3	3	3	3	3	3	3
4	4	4	4	4	4	4	4	4
5	5	5	5	5	5	5	5	5
6	6	6	6	6	6	6	6	6
7	7	7	7	7	7	7	7	7
8	8	8	8	8	8	8	8	8
9	9	9	9	9	9	9	9	9

## ► Data representation



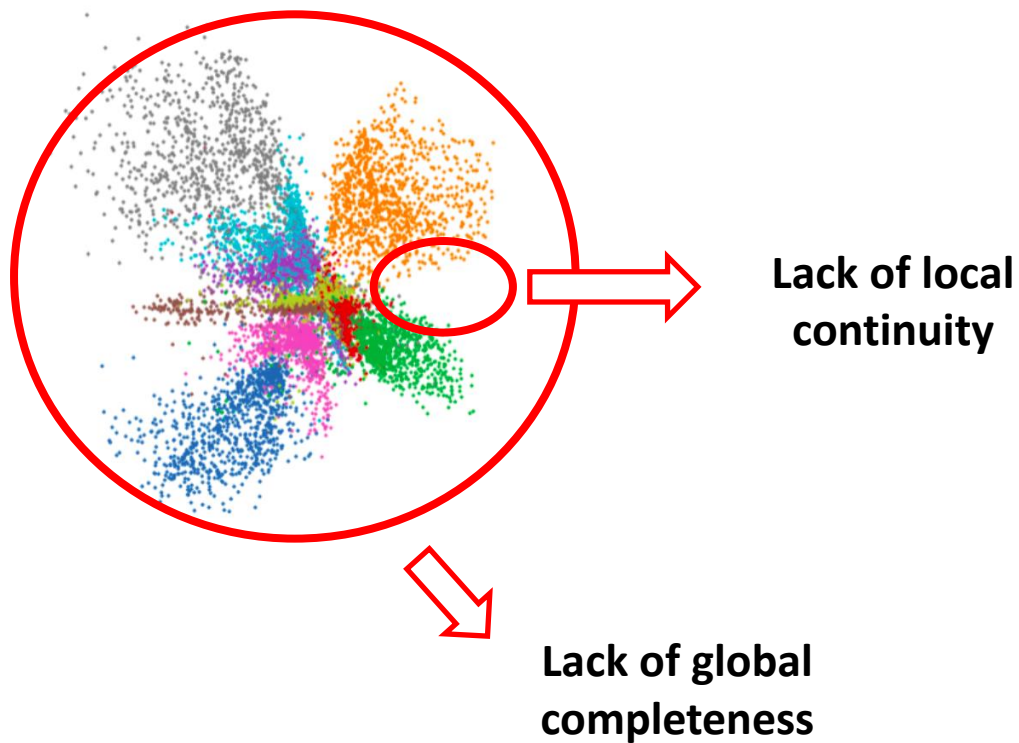
# Interest of auto-encoders

## ► Generative model



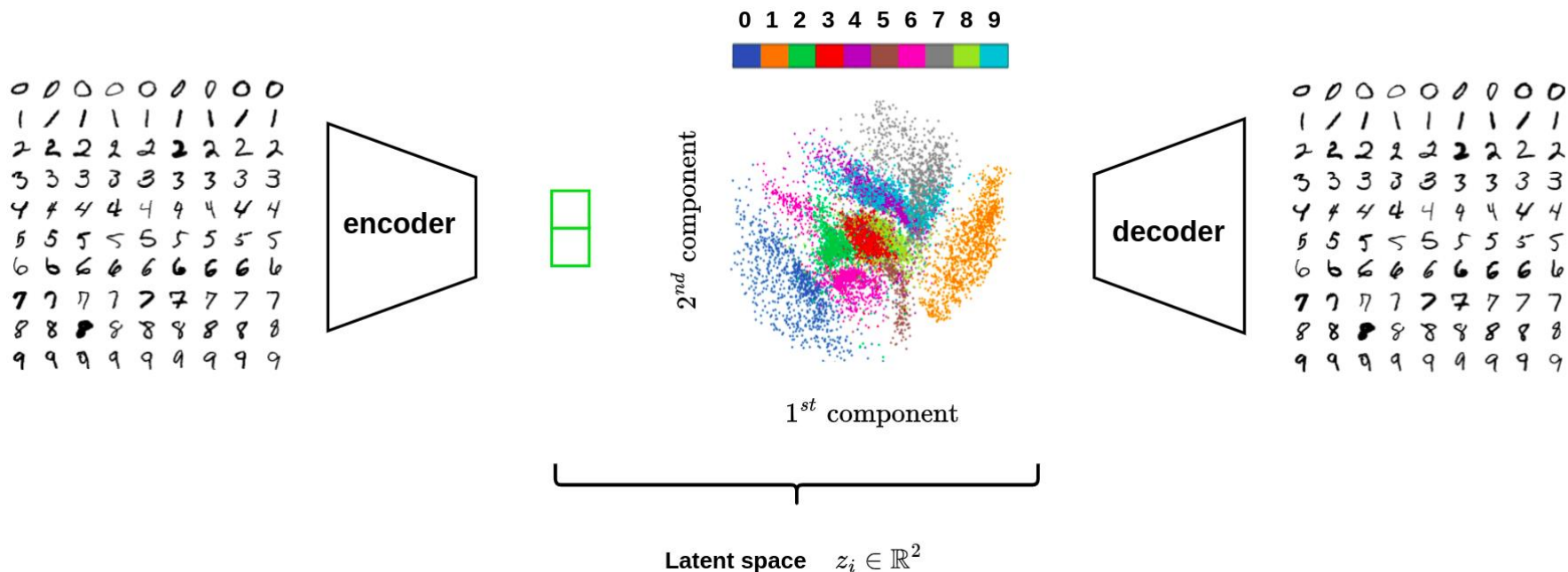
## Limitations

- ▶ Needs to better control the structure of the latent space

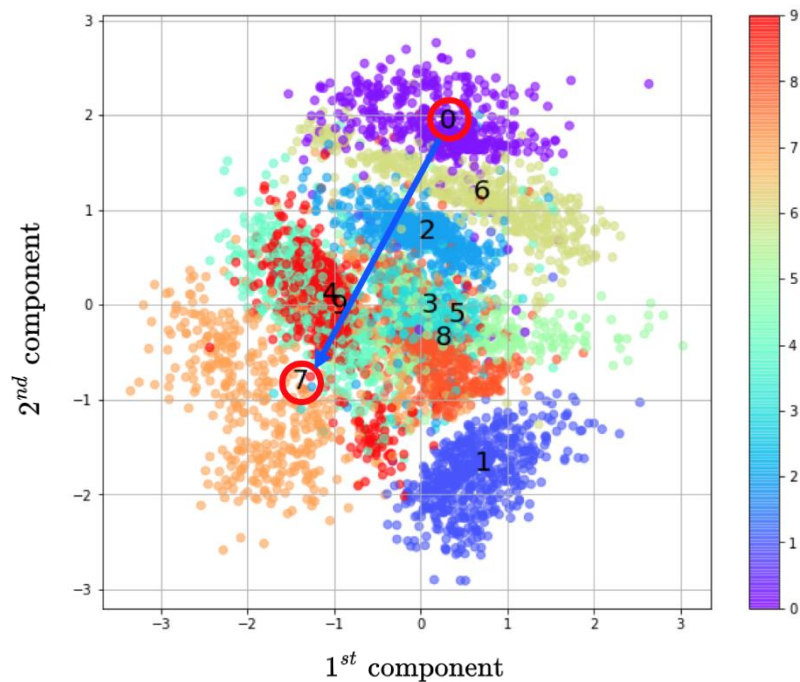


# Interest of auto-encoders

- ▶ Generative model with better properties thanks to *variational framework*

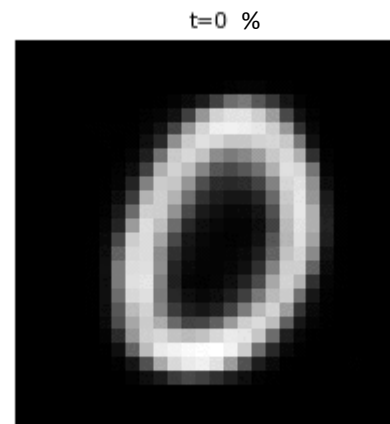


## ► Generative model with variational framework



Linear interpolation into the latent space

$$t \cdot z_0 + (1 - t) \cdot z_7, \quad 0 \leq t \leq 1$$





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# Variational autoencoders

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*All the mathematical details are given there !*

<https://creatis-myriad.github.io/tutorials/2022-09-12-tutorial-vae.html>

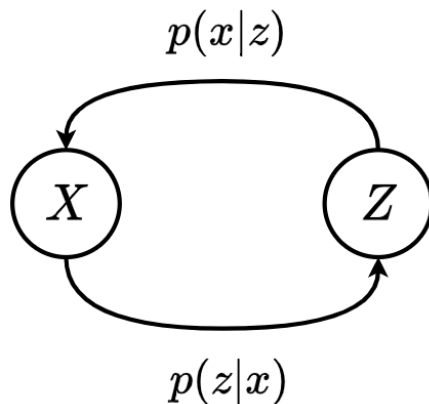
- ▶ **Enforcing a structured latent space**
  - **Through a probabilistic framework**
  - **By imposing continuity**
  - **By imposing completeness**

# Probabilistic framework

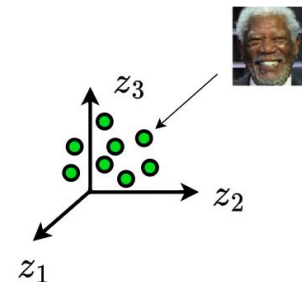
## ► Mathematical formulation



Observation  
variables



Hidden  
variables



Approximation of  $p(z|x)$  through a variational inference technique

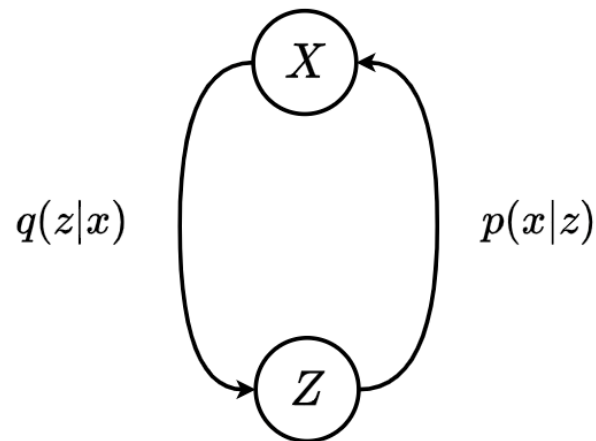
## ► Hypotheses

→  $q(z|x)$  is modeled by an axis-aligned Gaussian distribution

→  $q(z|x) = \mathcal{N}(\mu_x, \sigma_x) = \mathcal{N}(g(x), \text{diag}(h(x)))$

$$(g^*, h^*) = \arg \min_{(g, h)} D_{KL}(q(z|x) \parallel p(z|x))$$

$D_{KL}(\cdot \parallel \cdot)$  Kullback-Liebler divergence function



## ► Optimization process

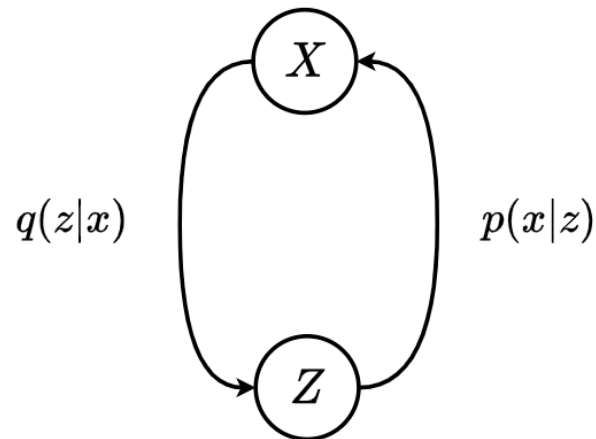
### → Maximization of the Evidence Lower Bound (ELBO)

$$\mathcal{L} = \mathbb{E}_{z \sim q_x} [\log(p(x|z))] - D_{KL}(q(z|x) \parallel p(z))$$

### → By exploiting gaussian assumption

$$p(x|z) = \mathcal{N}(f(z), cI)$$

$$\mathcal{L} \propto \mathbb{E}_{z \sim q_x} [-\alpha \|x - f(z)\|^2] - D_{KL}(q(z|x) \parallel p(z))$$



# Probabilistic framework

## ► Optimization process

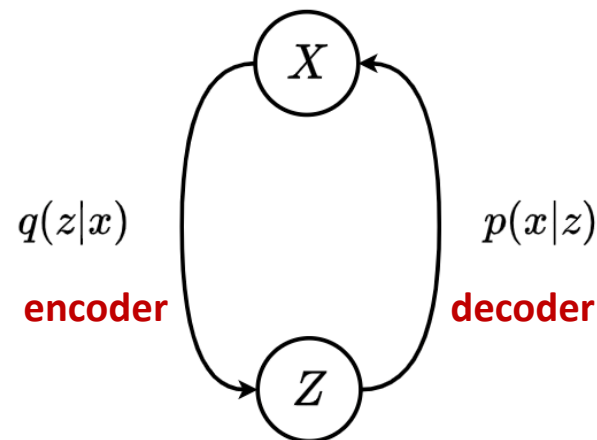
$$(f^*, g^*, h^*) = \arg \min_{(f, g, h)} (\mathbb{E}_{z \sim q_x} [\alpha \|x - f(z)\|^2] + D_{KL}(q(z|x) \parallel p(z)))$$

## ► Deep learning loss function

$$\text{loss} = \alpha \|x - f(z)\|^2 + D_{KL}(\mathcal{N}(g(x), \text{diag}(h(x))), \mathcal{N}(0, I))$$

→  $g(\cdot)$  and  $h(\cdot)$  are modeled through an encoder

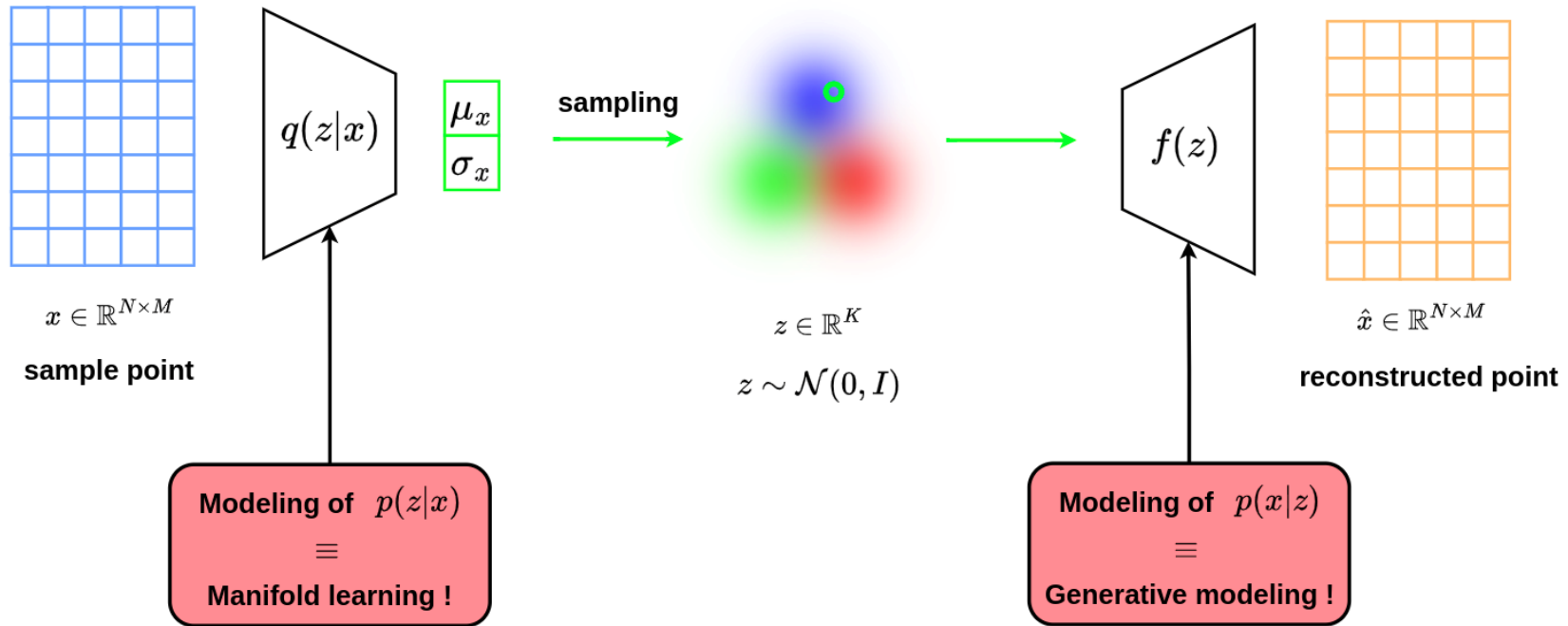
→  $f(\cdot)$  is modeled through a decoder



# Probabilistic framework

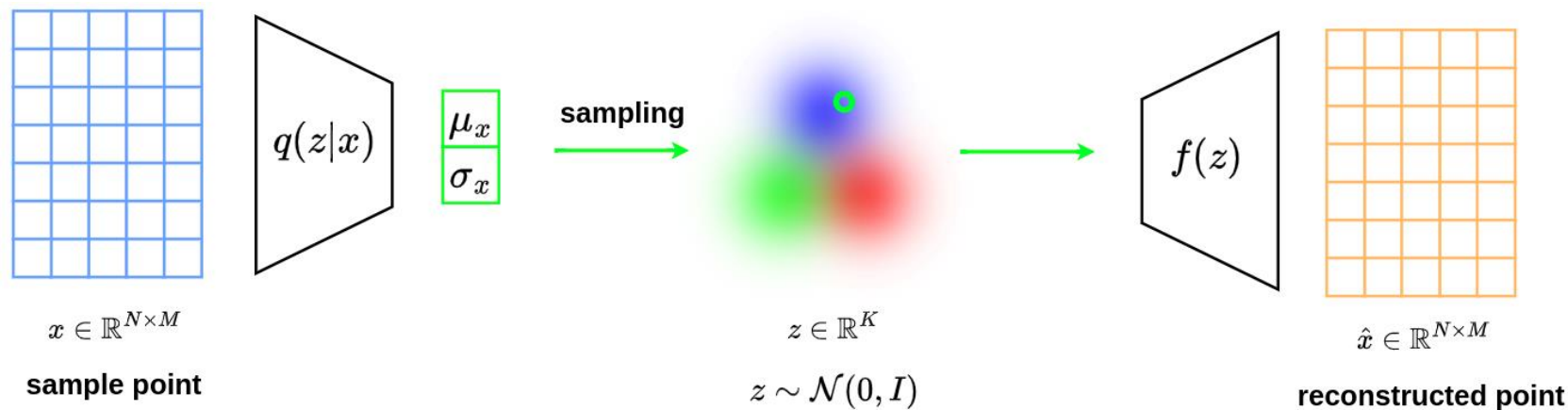
## ► Loss interpretation

$$\text{loss} = D_{KL}(\mathcal{N}(g(x), \text{diag}(h(x))), \mathcal{N}(0, I)) + \alpha \|x - f(z)\|^2$$



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$$\text{loss} = D_{KL}(\mathcal{N}(g(x), \text{diag}(h(x))), \mathcal{N}(0, I)) + \alpha \|x - f(z)\|^2$$

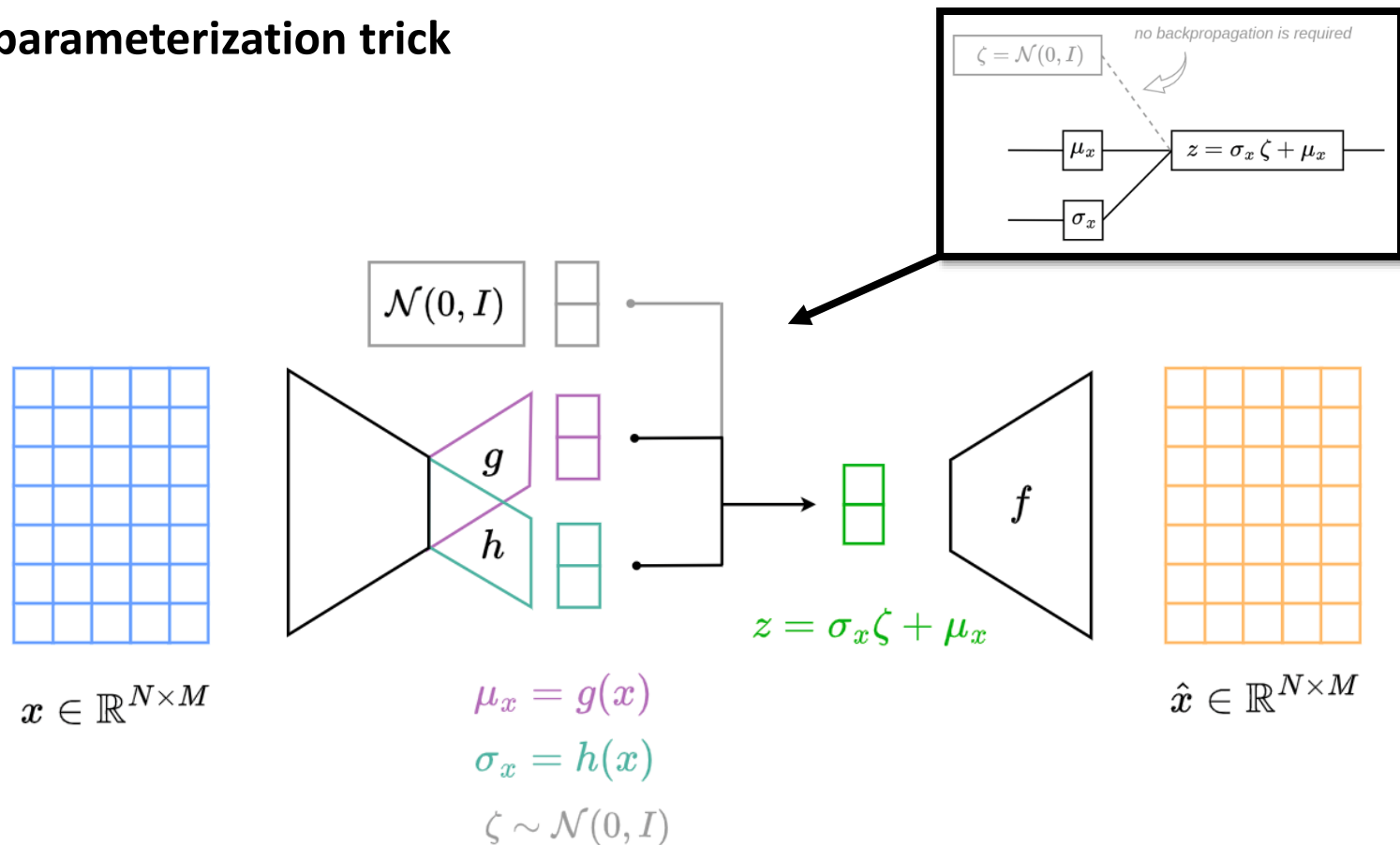


→  $\mathcal{N}(g(x), h(x))$  imposes local **continuity**

→  $\mathcal{N}(\cdot, \mathcal{N}(0, I))$  imposes global **completeness**



## ► Reparameterization trick



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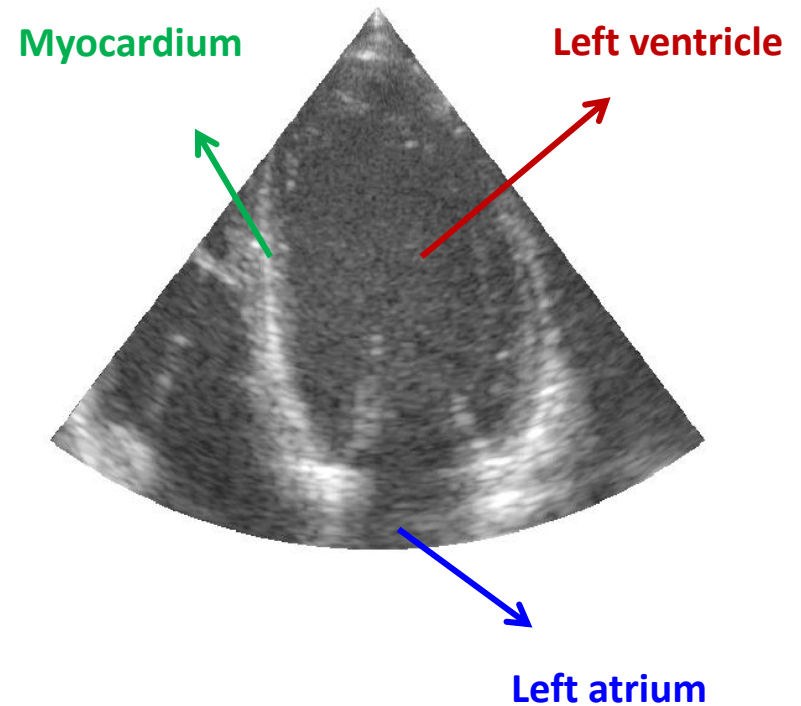
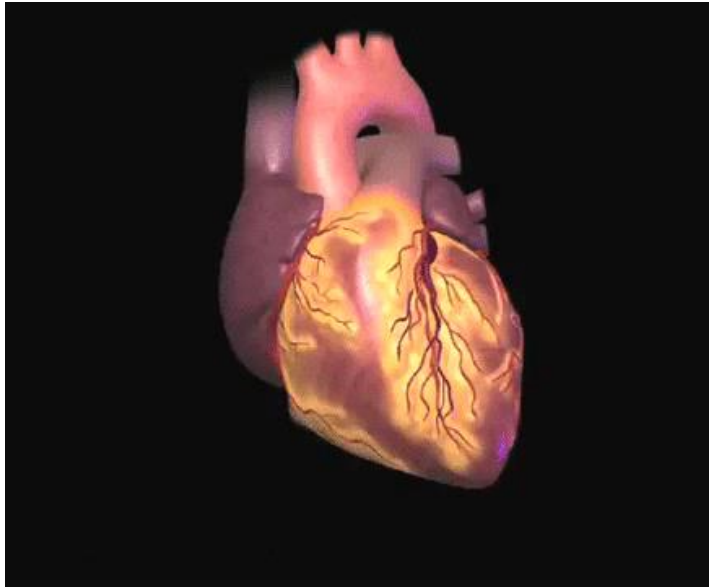
# Practical applications

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**The obsession is to master the latent space !!!**

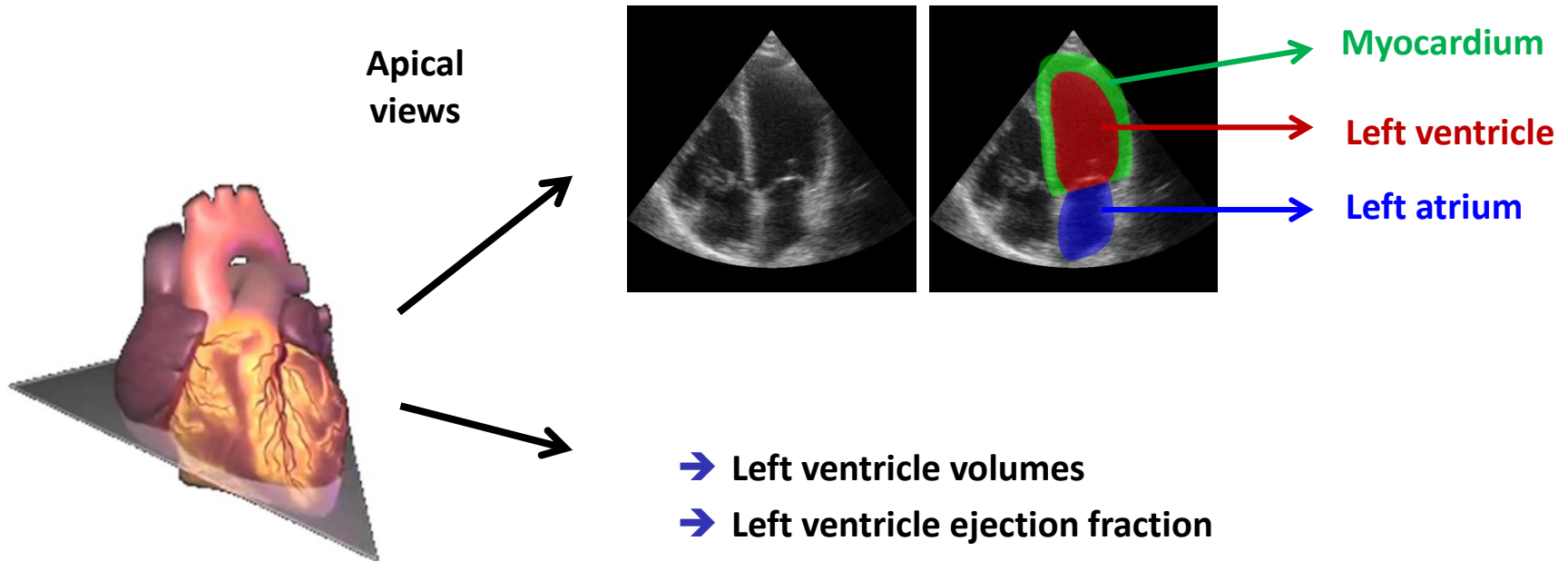
## Needs for accurate and robust segmentation of cardiac structures

- ▶ Quantification of clinical indices from echocardiographic images



# Needs for accurate and robust segmentation of cardiac structures

## ► Anatomical clinical indices



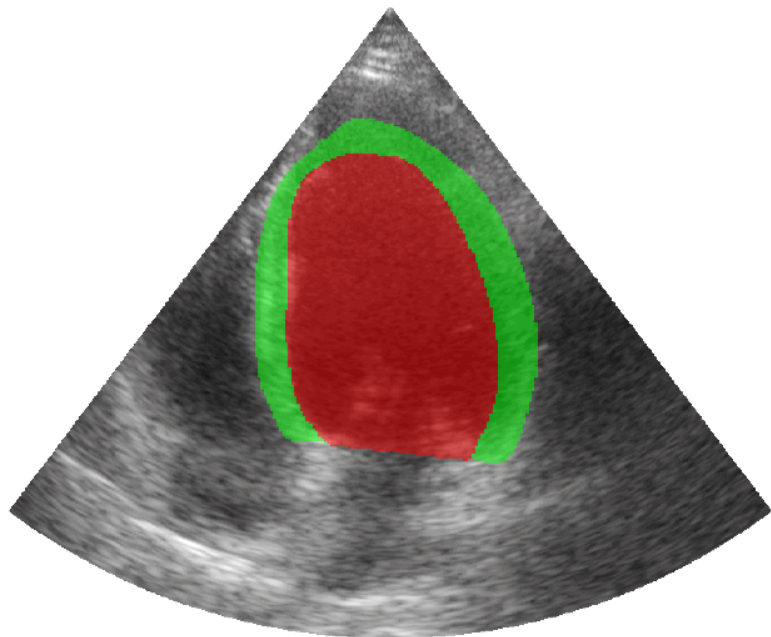
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# How to guarantee temporal consistency ?

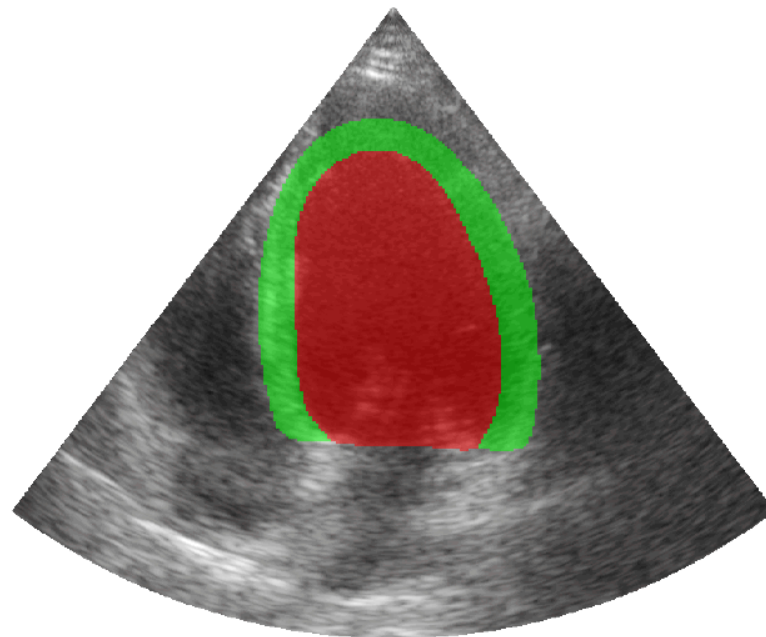
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- ▶ Quantification of clinical indices from echocardiographic images

What we have with a 2D U-Net



What we want



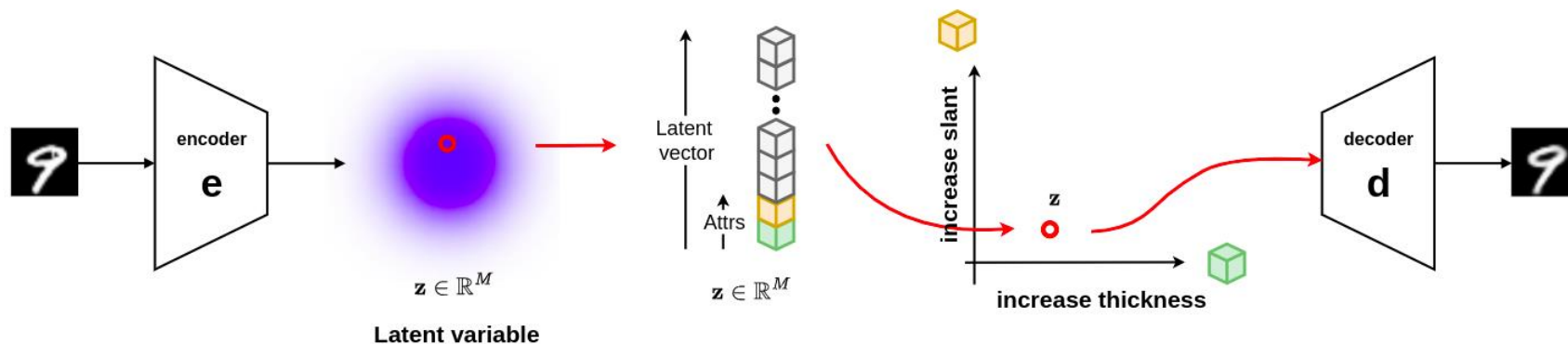
► AR-VAE: attribute-based regularization of VAE latent space

[Pati, Neural Comp. Appli., 2021]

- Generation of structured latent space

- ➔ Specific continuous-valued attributes forced to be encoded along specific dimensions

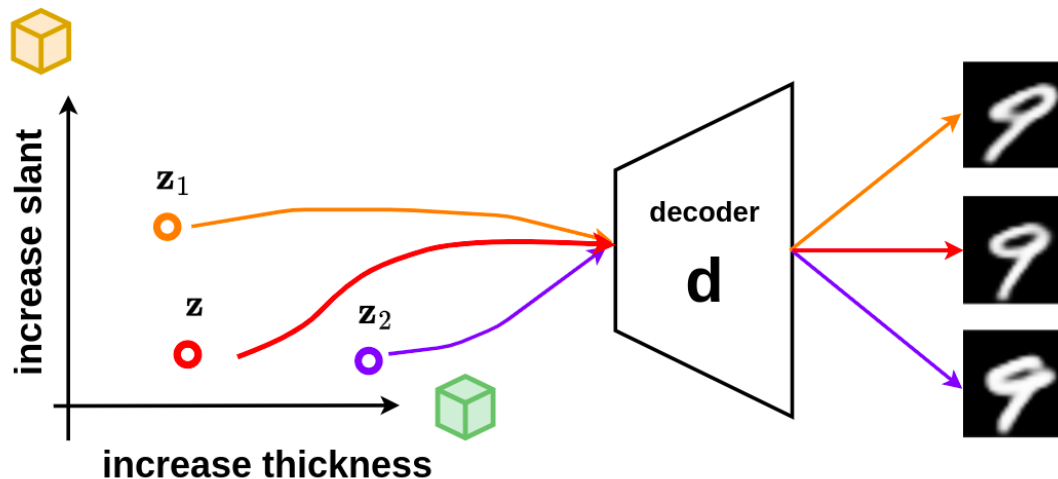
- ➔  $Loss = VAE\ loss + Attribute\ Regularisation\ Loss$



► AR-VAE: attribute-based regularization of VAE latent space

[Pati, Neural Comp. Appli., 2021]

- Sampling of the structured latent space





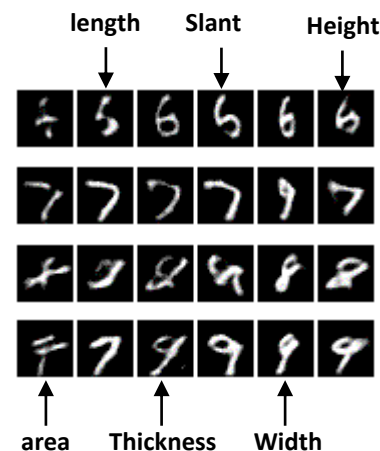
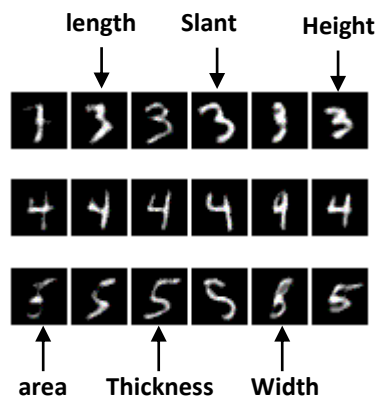
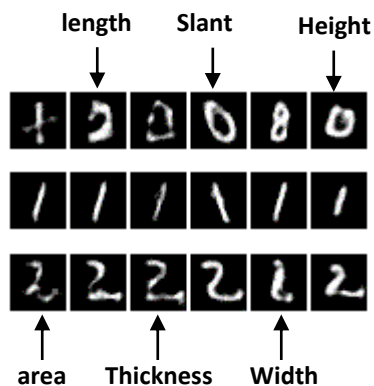
## ▶ AR-VAE: attribute-based regularization of VAE latent space

[Pati, Neural Comp. Appli., 2021]

- Sampling of the structured latent space

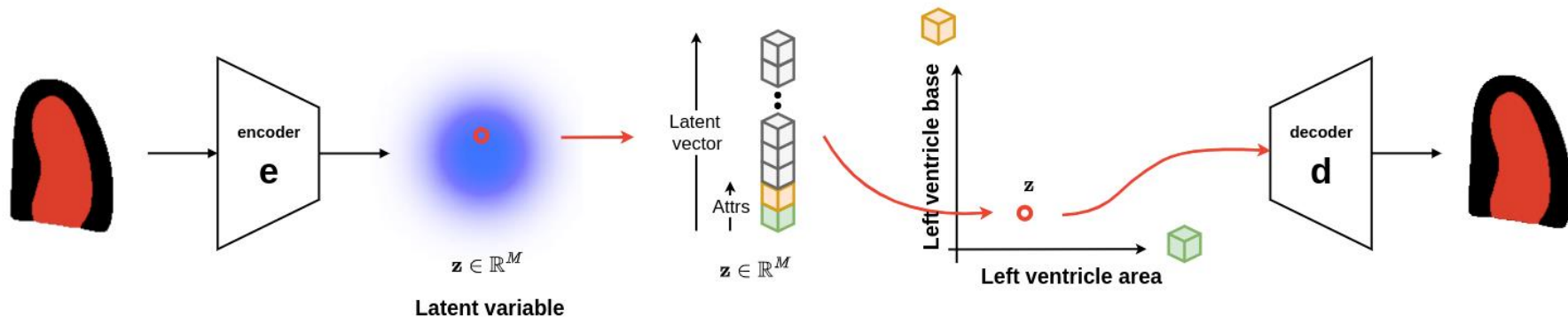
- ➔ Specific attribute (from left to right): area, length, thickness, slant, width, height

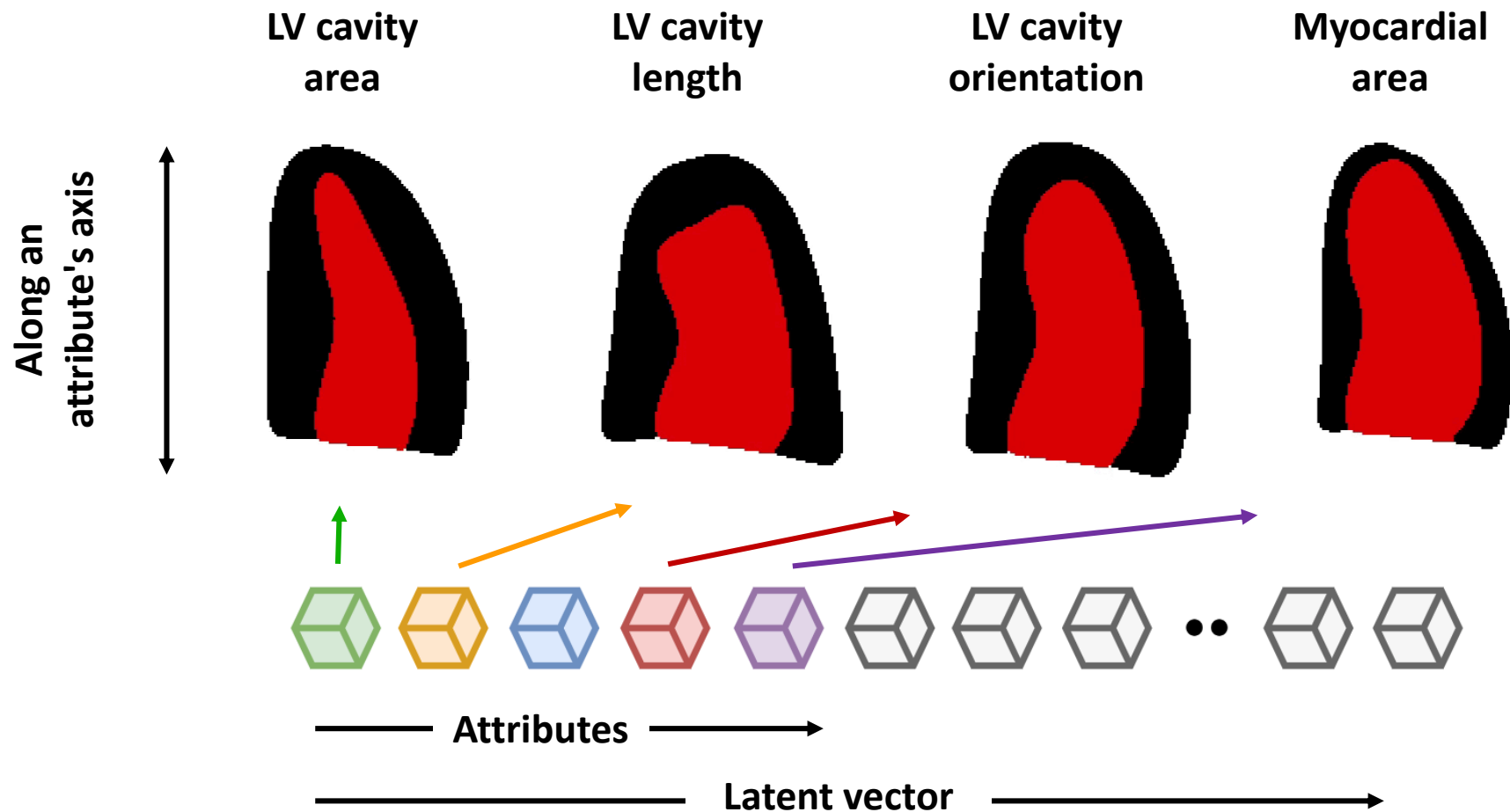
- ➔ Each column corresponds to traversal along a regularized dimension



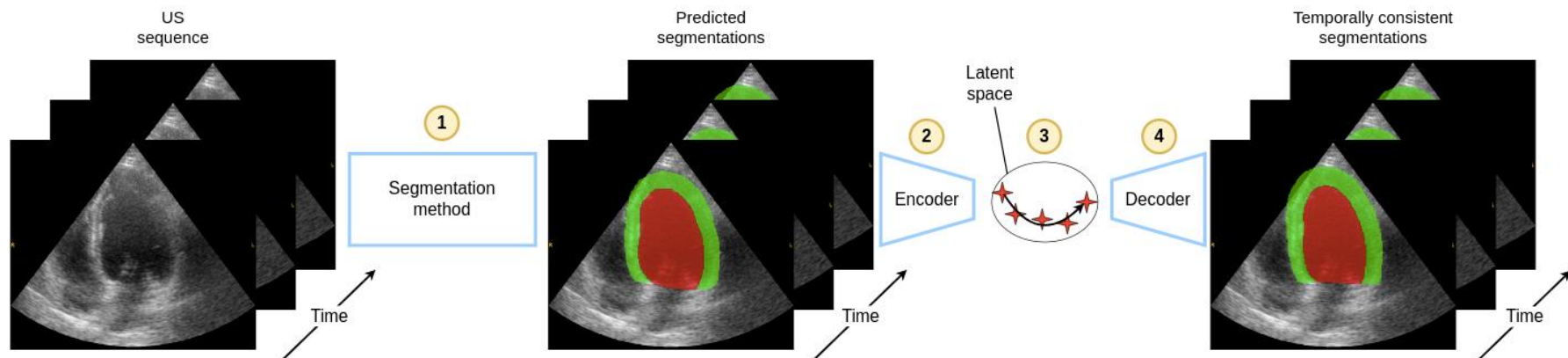
## ► Application to the description of the cardiac shapes

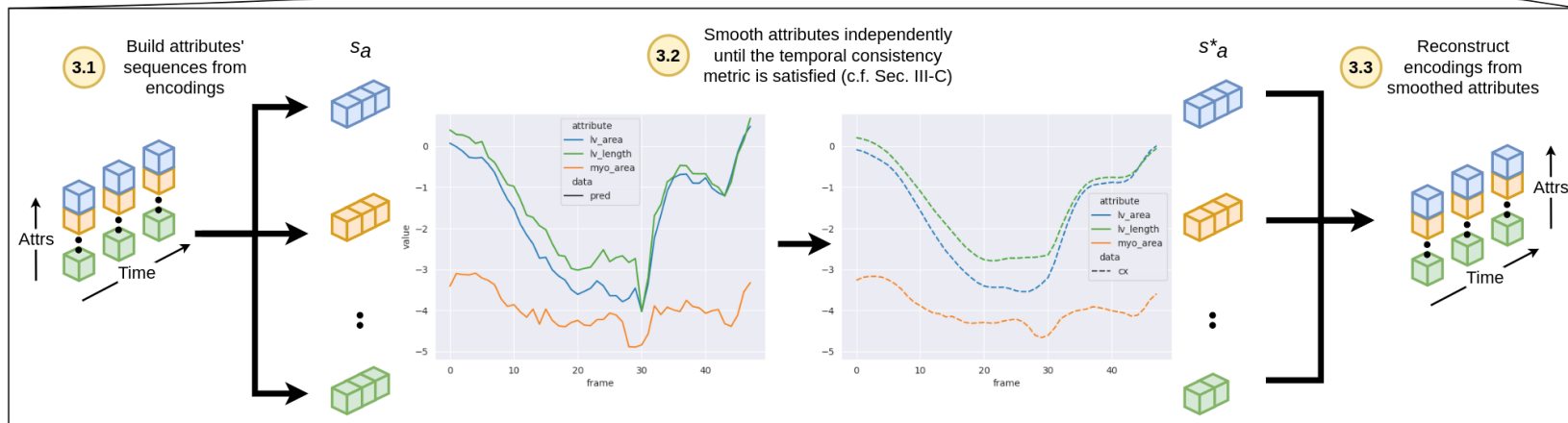
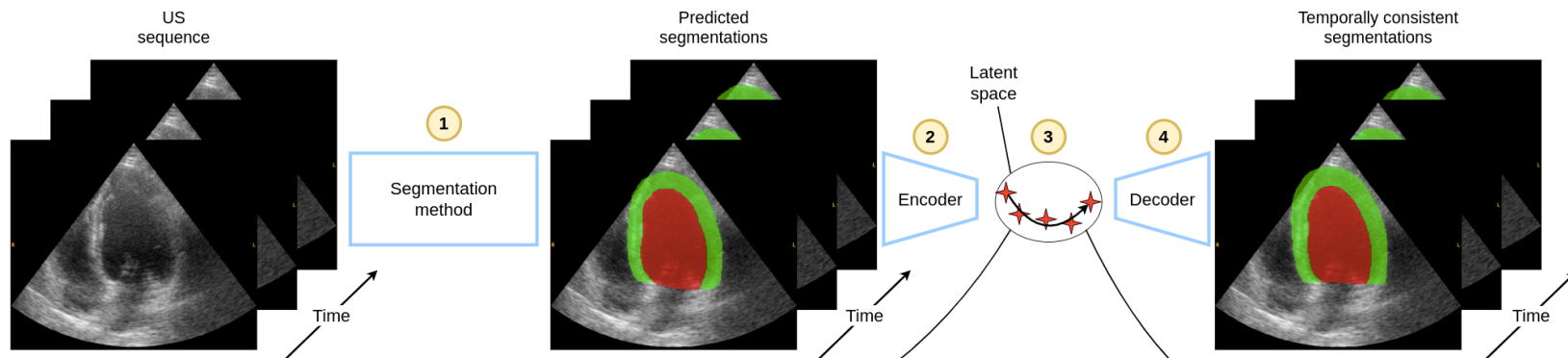
- Generation of structured latent space according to the following attributes
  - ➔ Left ventricle (LV) cavity: area, length, basal width, orientation
  - ➔ Myocardial area
  - ➔ Epicardial center





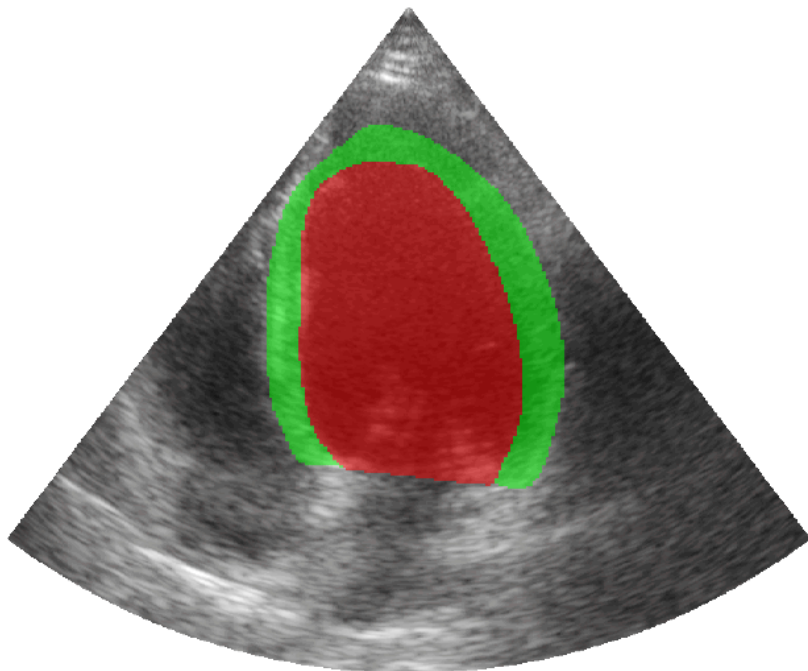
## ► Proposed temporal pipeline



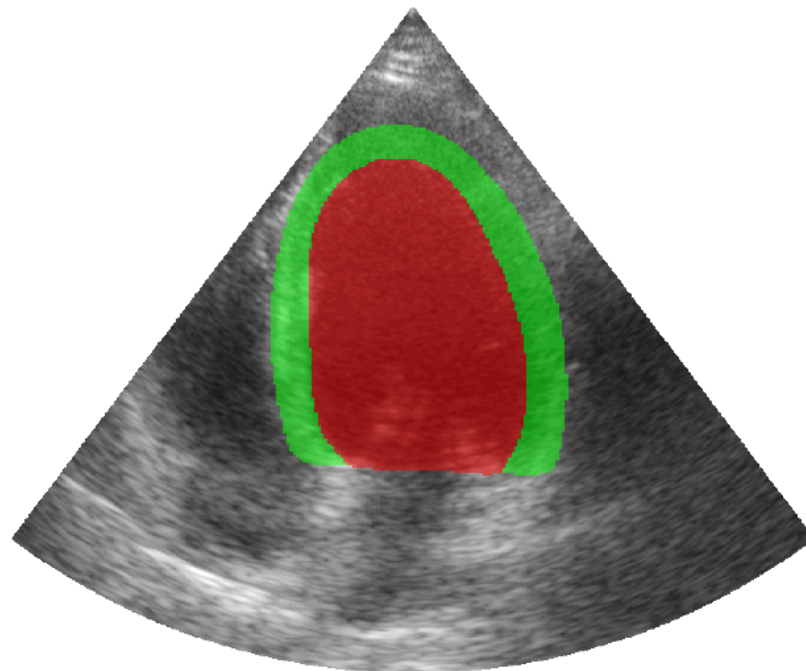


► Some post-processing examples

Original U-Net

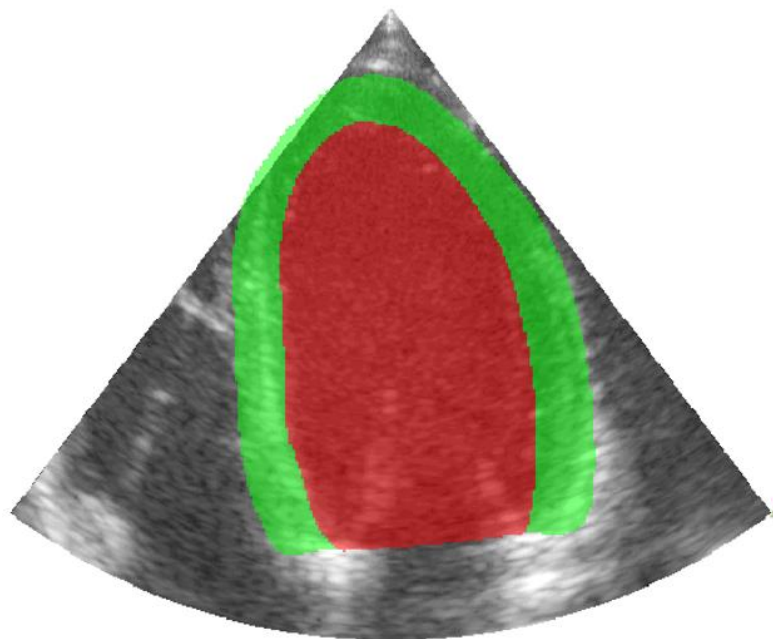


Post-processed U-Net

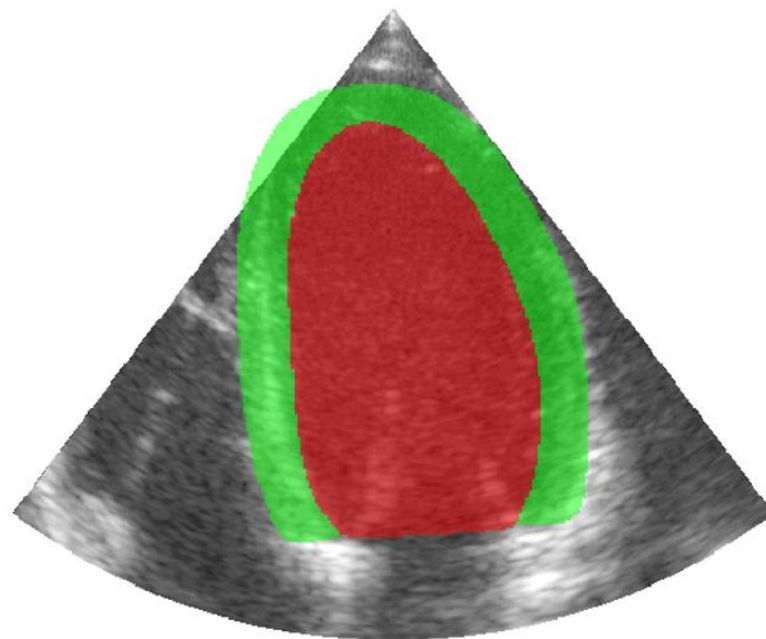


► Some post-processing examples

Original U-Net



Post-processed U-Net



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**To conclude**

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## To conclude

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▶ **VAEs can be used effectively in medical imaging**

- **Guarantee anatomical coherence** ✓
- **Guarantee temporal consistency** ✓
- **Estimation uncertainty for image segmentation** ✓
- **Generative interest limited to simple distribution**

▶ **Useful tool for characterizing populations**

- **Need to properly structure the learned latent space**
- **Need to work on relatively large cohorts**

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# Appendix

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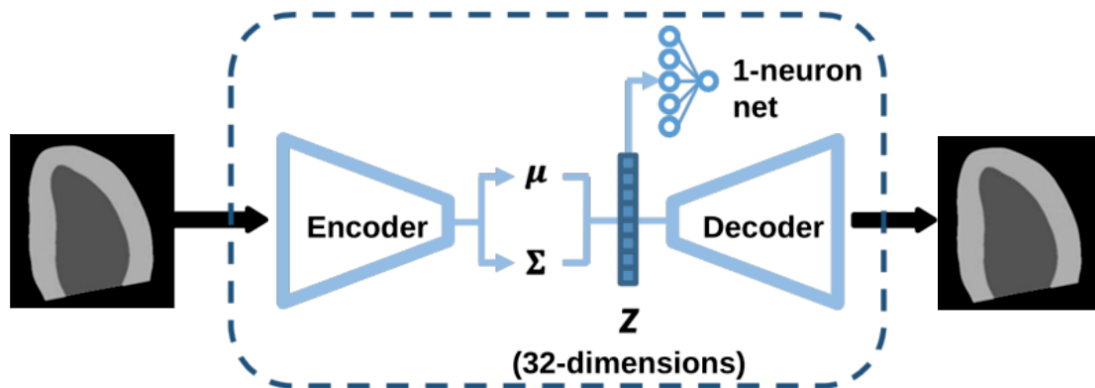
# How to guarantee the anatomical coherence ?

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► **Constrained Variational Auto Encoder**

- Approximation of a latent space with local linear properties

Use of a 1-neuron net to reinforce the linearity of the latent space

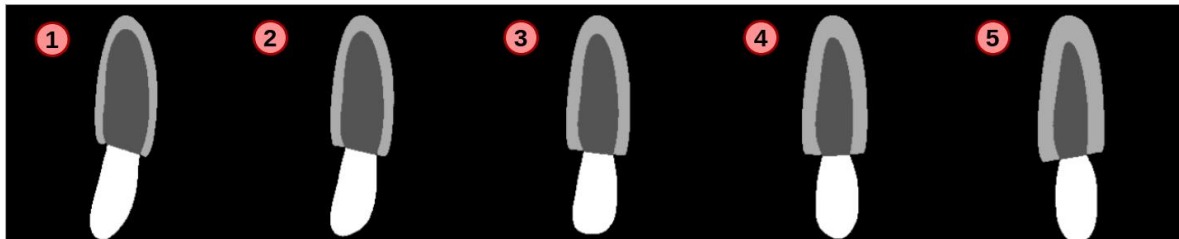
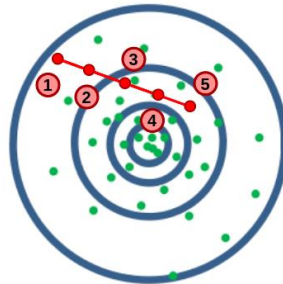


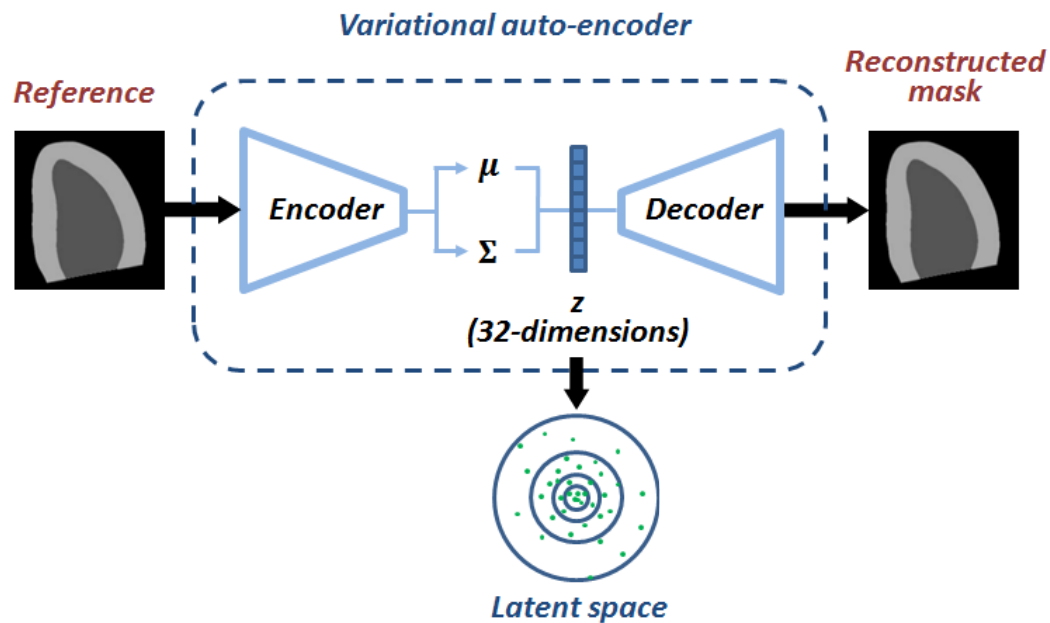
## ► Constrained Variational Auto Encoder

- Approximation of a latent space with local linear properties

→ Linear interpolation in the latent space makes sense

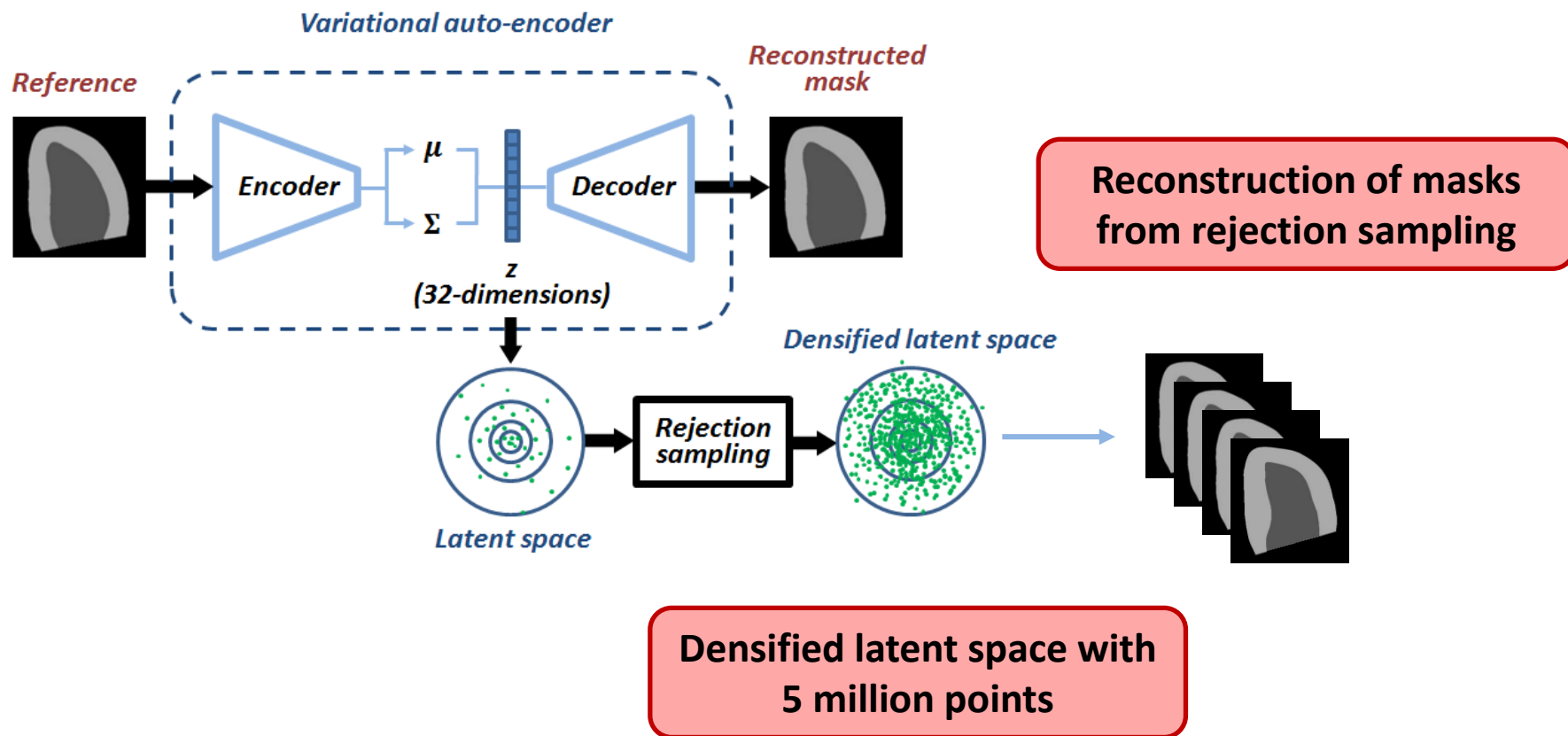
*Constrained  
latent space*





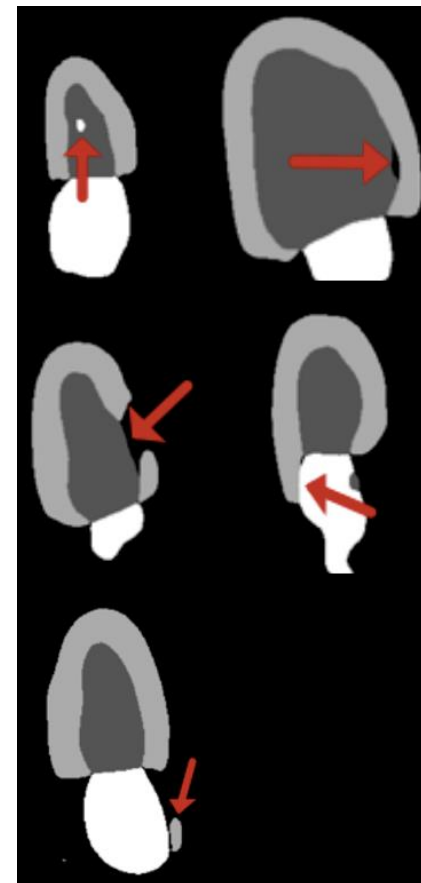
Efficient encoding of anatomical shapes in a latent space

# Cardiac segmentation with strong anatomical guarantees [Painchaud, IEEE TMI, 2020]



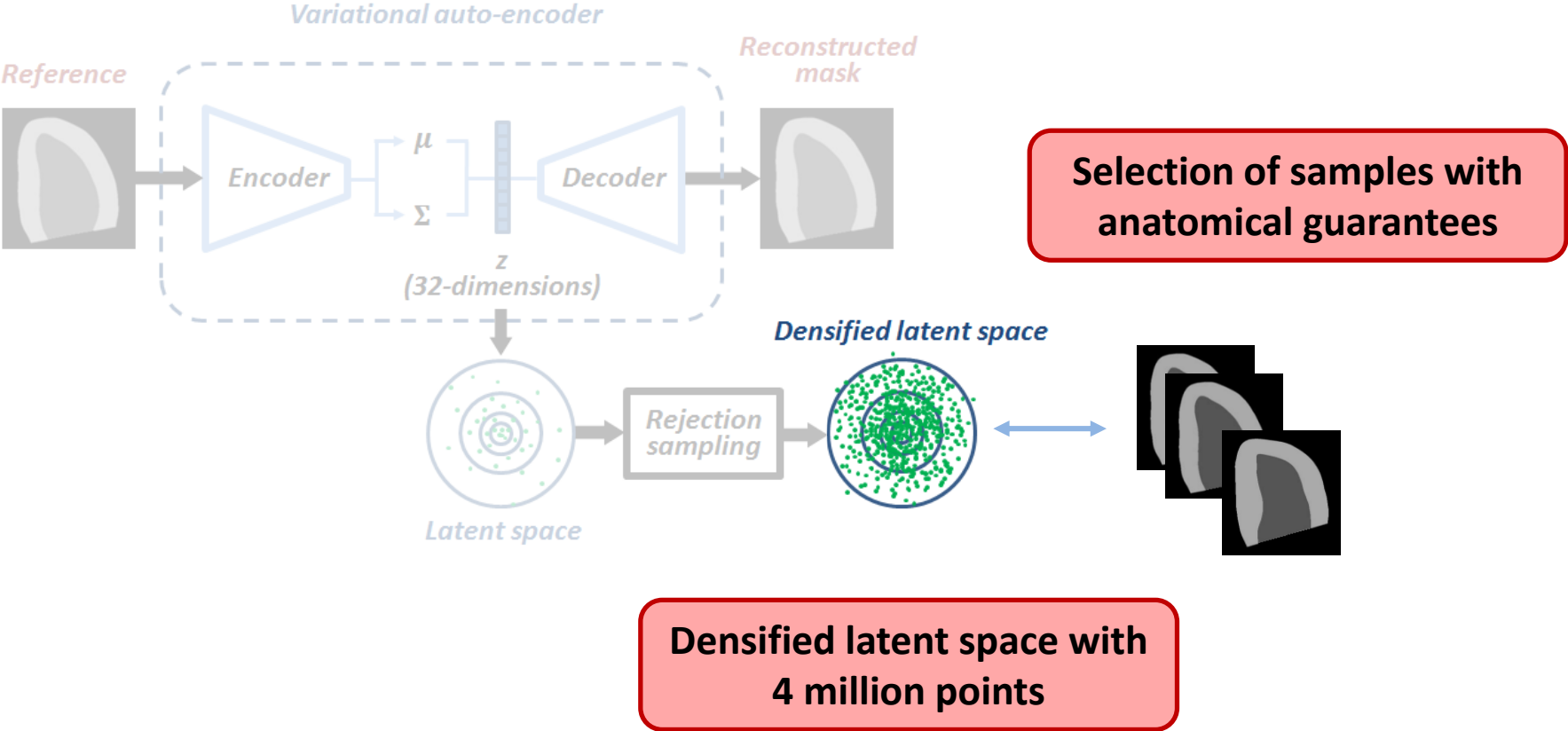
## ► Definition of 12 anatomical metrics

- (3 criteria) hole(s) in the LV, RV or LA
- (2 criteria) hole(s) between LV and MYO or between LV and LA
- (3 criteria) presence of more than one LV, MYO or LA
- (2 criteria) size of the area by which the LV touches the background or the MYO touches the LA
- (1 criterion) ratio between the minimal and maximal thickness of the MYO
- (1 criterion) ratio between the width of the LV and the average thickness of the MYO





# Cardiac segmentation with strong anatomical guarantees [Painchaud, IEEE TMI, 2020]

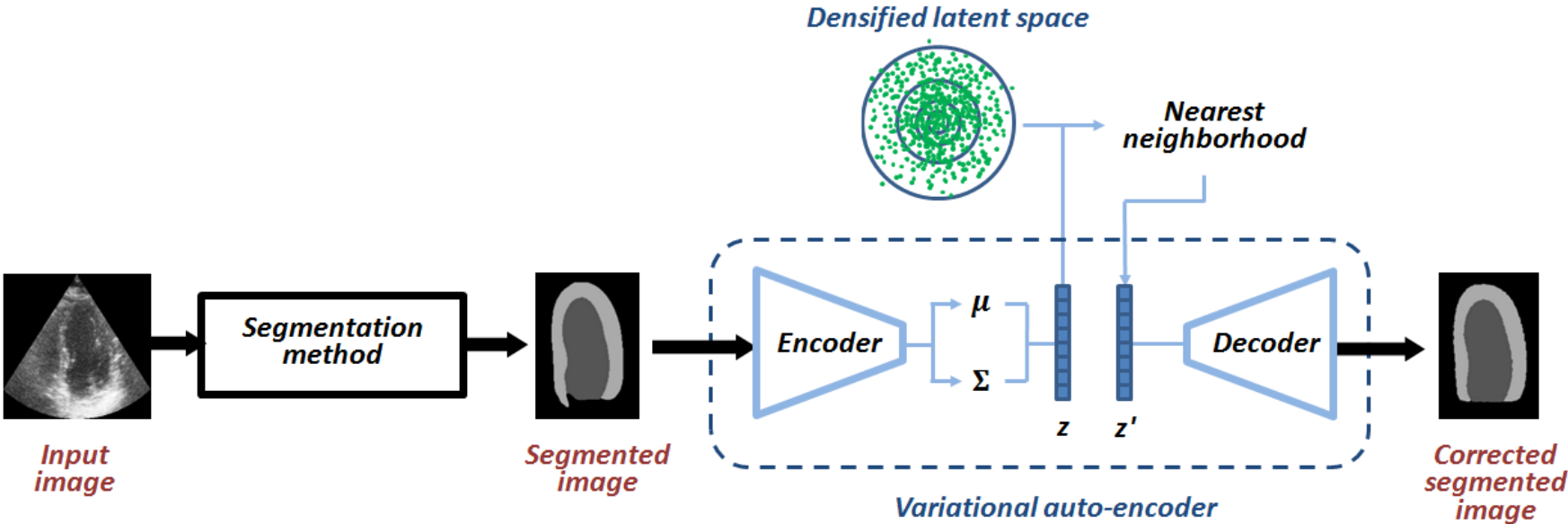


# Cardiac segmentation with strong anatomical guarantees [Painchaud, IEEE TMI, 2020]

Correction of segmentation to guarantee the plausibility of anatomical shapes



Almost same accuracy as the original methods but with correct anatomical shapes

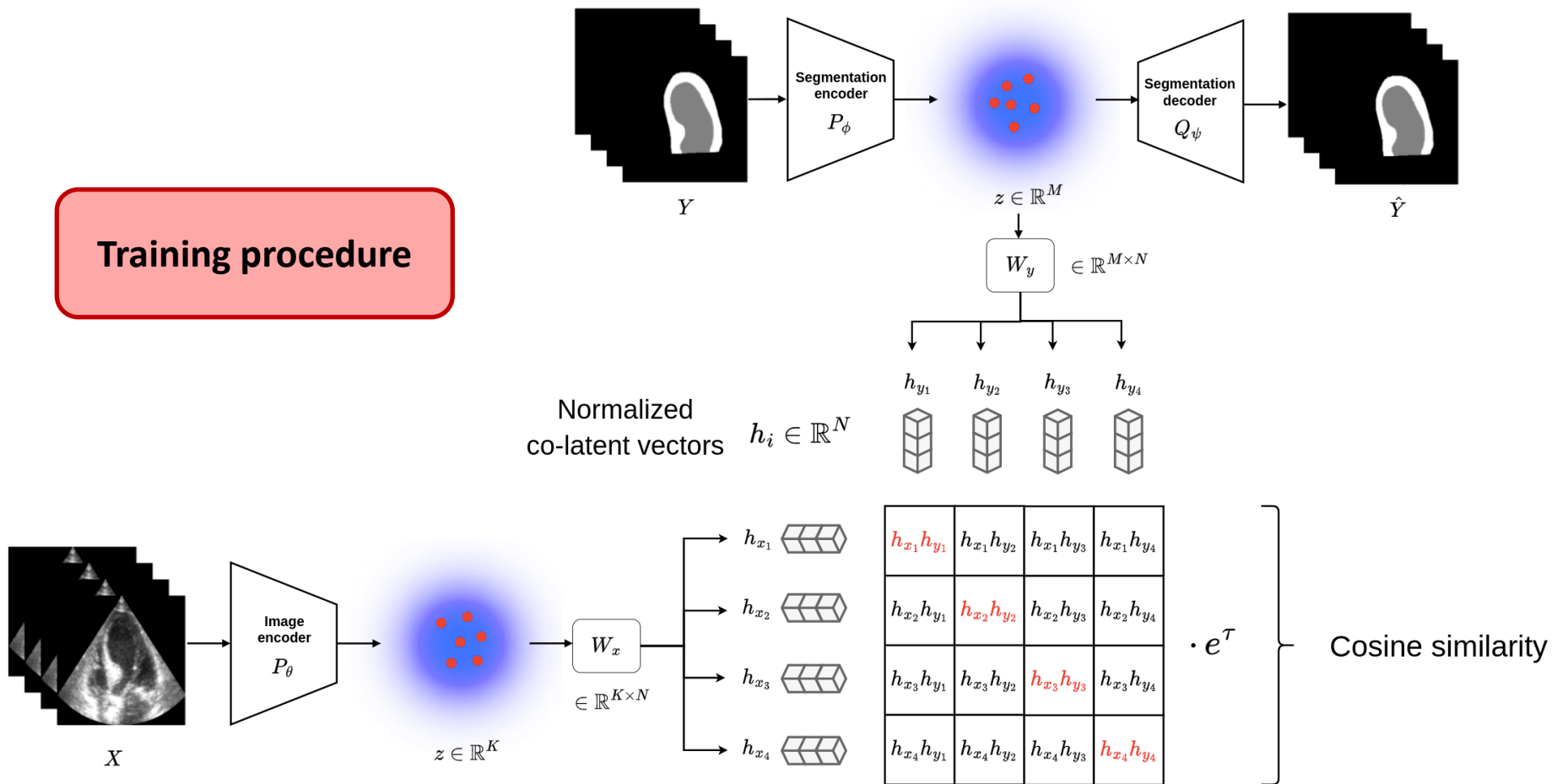


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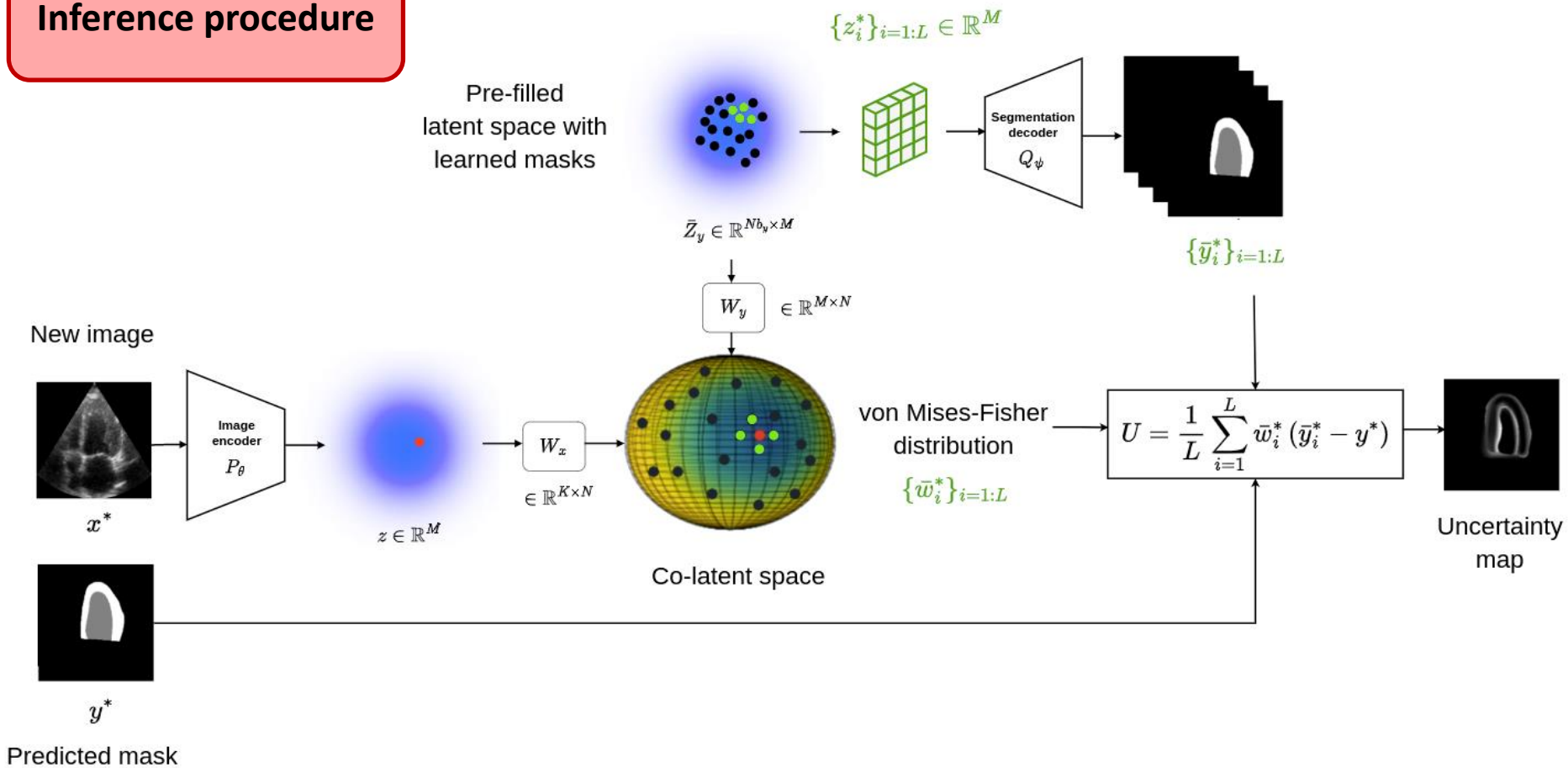
# Uncertainty estimation for cardiac image segmentation

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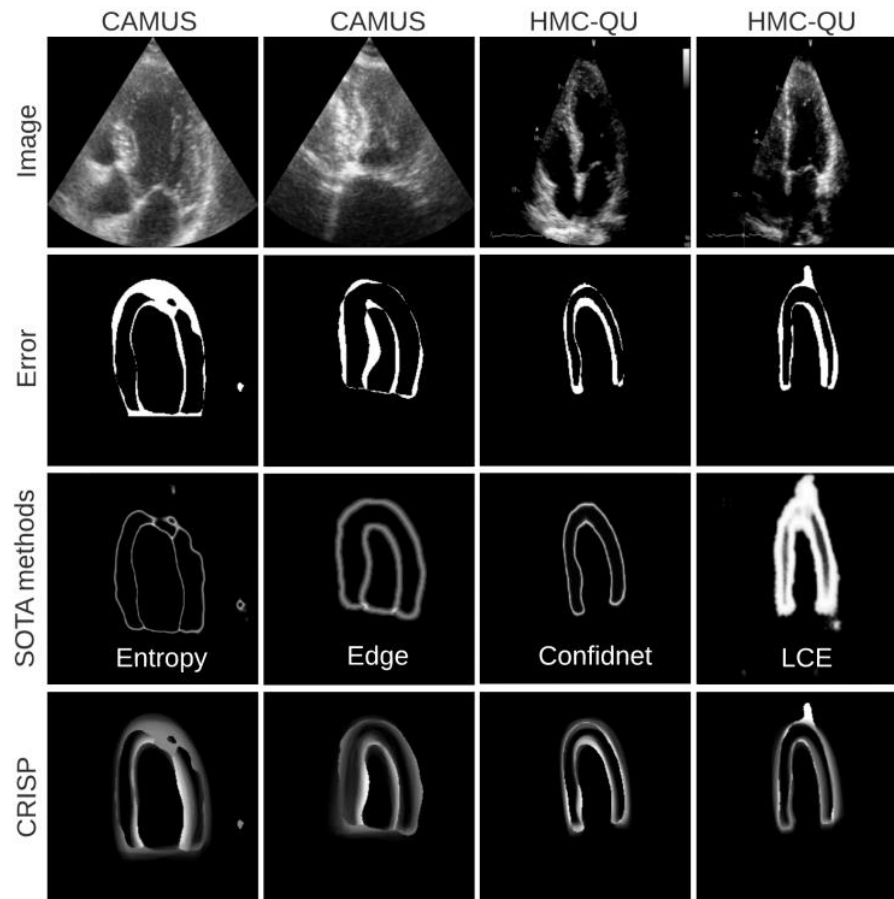
**Training procedure**



**Inference procedure**



## ► Uncertainty results



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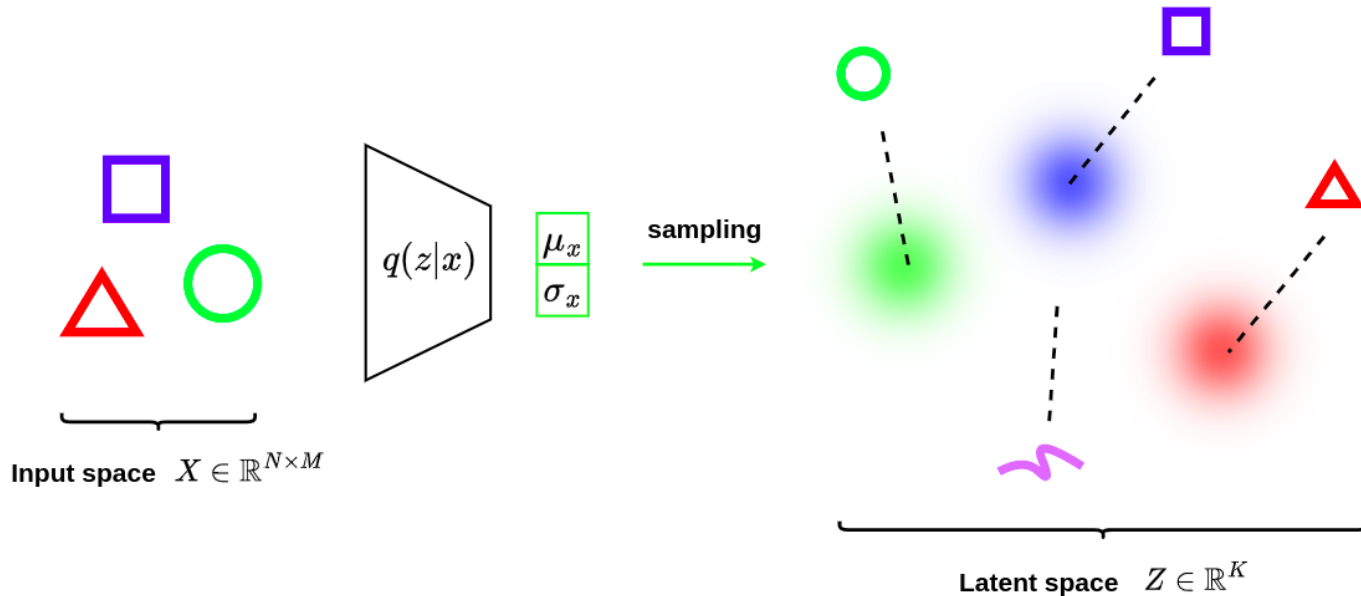
# Misc

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# Probabilistic framework

## ► Continuity

$$\mathcal{N}(g(x), h(x))$$

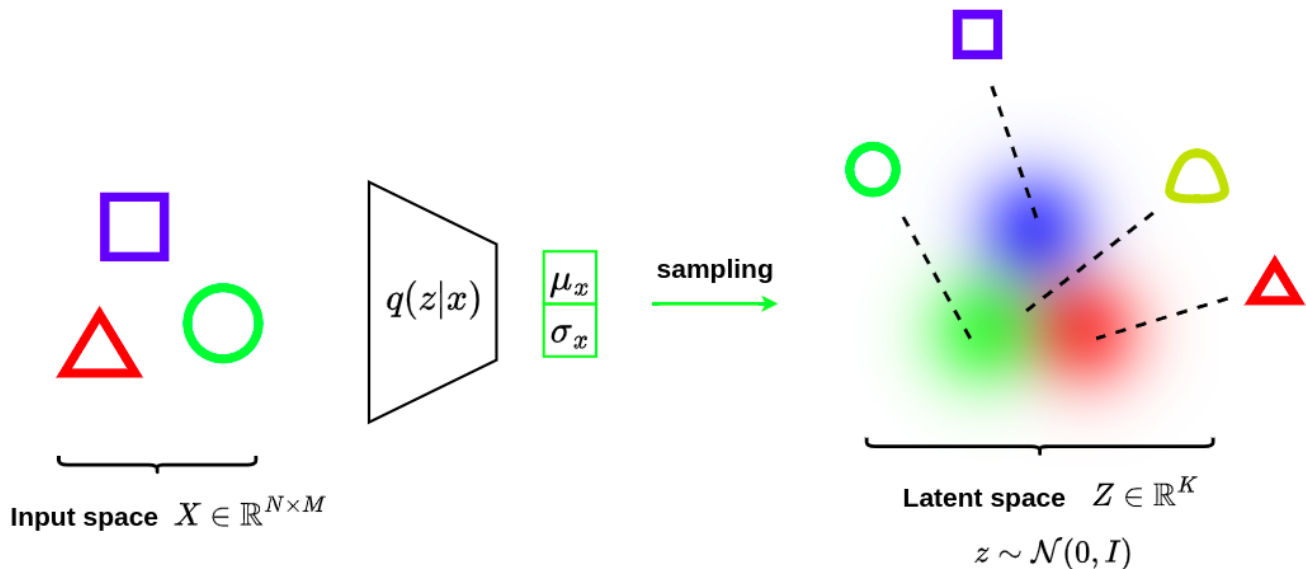




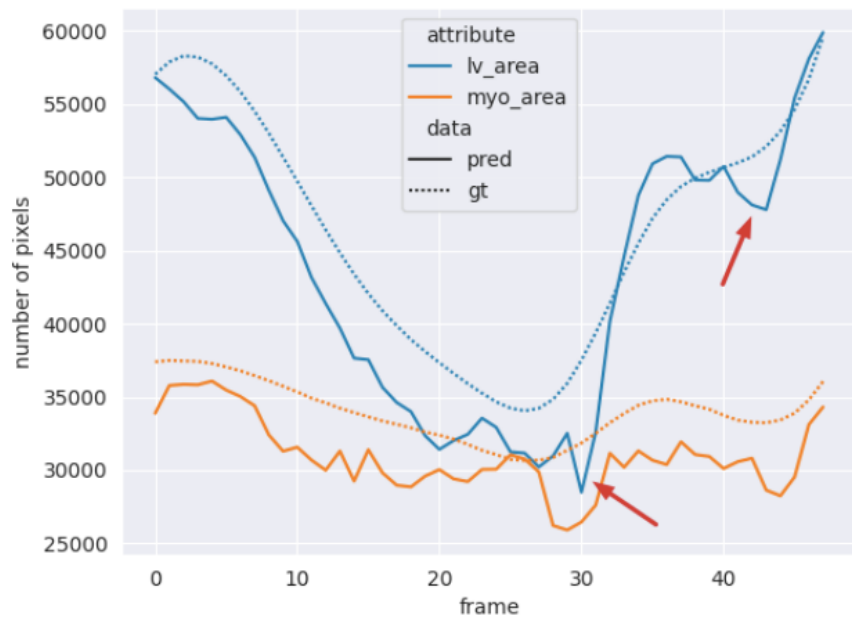
# Probabilistic framework

## ► Completeness

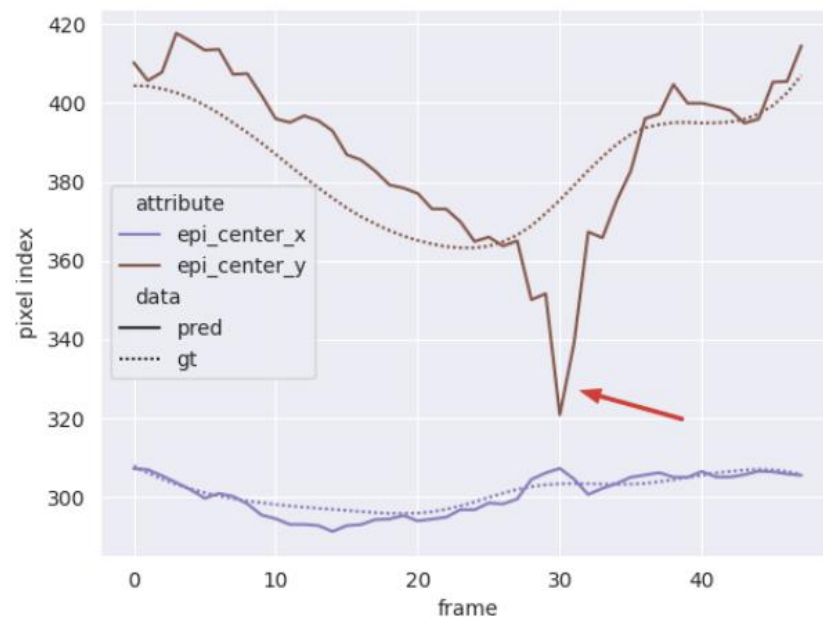
$$\mathcal{N}(\cdot, \mathcal{N}(0, I))$$



## ► Temporal inconsistency detection from the latent space



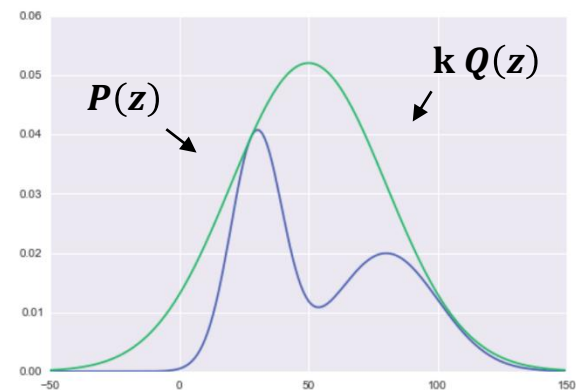
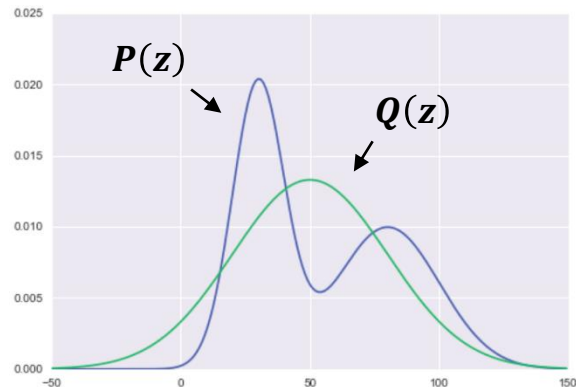
**Choppy contraction/dilation of the LV cavity**



**Abrupt vertical shifts of the cardiac shape**

## ► Rejection sampling

- Targeted distribution  $P(z)$ 
  - ➔ Parzen window technique
- Proposed distribution  $Q(z)$
  
- Constrain  $kQ(z) > P(z)$ 
  - ➔ Automatic choice of  $k$



## ► Rejection sampling

- $z \sim Q(z)$
- $u \sim \text{Unif}(0, kQ(z))$
- Computation of  $P(z)$ 
  - If  $u \leq P(z)$  then keep  $z$
  - If  $u > P(z)$  then reject  $z$

