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# Background

• In single particle electron microscopy, we are interested in experimentally determining the structure of a protein Statistical methods assume that the many observations (2D projections) of the protein arise from either a single structure or from a discrete mixture of structures

• In 2D classification, the protein views are also grouped into discrete clusters

• These conformations are confounded by orientation in the collected images

Motivation: learn generative models that are orientation invariant. That is, we wish to structure our generative model to enable structured inference on the unobserved orientations.

# Problem

Learn a generative model of 2-D signals (y) in which the content is described by a set of unstructured, continuous latent variables, z. That is

$$z \sim N(0, I),$$
  
 $v \sim f(z).$ 

However, we observe noisy images that are corrupted with translation and rotation (in EM, particles are randomly oriented), but we do not wish to encode these features into z.



Parameterize the generative model as a function of the spatial coordinates, i.e. the log probability of an image, y, given latent variables, z, and spatial coordinates, x, is

$$\log p(y|z) = \sum_{i=1}^n \log p_g(y^i|x^i, z)$$

This allows us to naturally model rotation and translation as transformations of the spatial coordinates. Let  $R(\theta)$  be the rotation matrix for angle  $\theta$  and  $\Delta x$  be the translation, then the log probability of the image is given by

$$\log p_g(y|z,\theta,\Delta x) = \sum_{i=1}^n \log p_g(y^i|x^i R(\theta) + \Delta x, z)$$

We then perform approximate inference on z,  $\theta$ , and  $\Delta x$  using a variational autoencoder framework.

# translation and rotation with spatial-VAE



Modeling galaxies from the Sloan Digital Sky Survey

- variables encoding content from rotation and translation

• The general framework extends to higher dimension signals and other coordinate transformations

## Conclusions

• Spatial-VAE enables learning generative models of images that separate latent

• The framework learns continuous conformations of proteins and galaxies in 2D