## Second year review

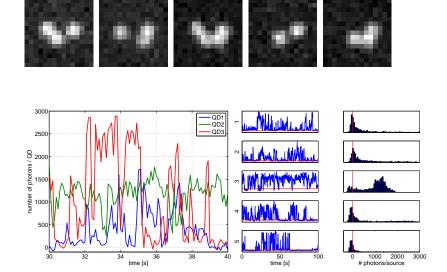
Ondřej Mandula

9th September 2011

### Overview

- 1. Introduction
- 2. Model comparison
- 3. Theoretical limits of the LM
- 4. Out of focus PSF
- 5. Future work

## Introduction: Quantum Dots and LM

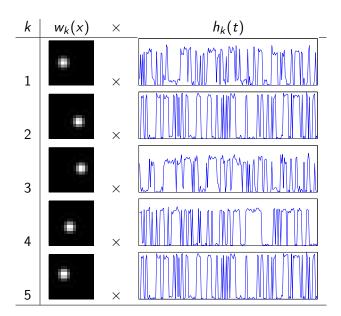


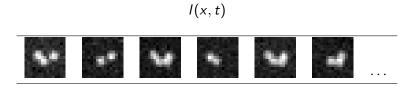
model of one source

$$s_k(x, t) = w_k(x)h_k(t)$$

model of an image

$$I(x,t) = \sum_{k=1}^{N} s_k(x,t)$$
$$= \sum_{k=1}^{N} w_k(x)h_k(t)$$





time  $\rightarrow$ 

matrix form - factorisation problem

$$I(x, t) = W(x)H(t)$$

relaxed model

$$E[I(x,t)] = W(x)H(t)$$

non-negativity constraints

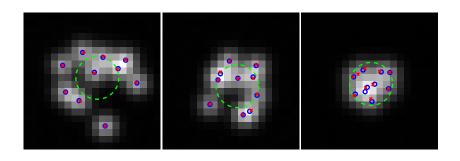
$$l \ge 0, \ W \ge 0, \ H \ge 0$$

## Algorithms

- Maximum a posteriori fitting (MAP)
  [Harrington et al., 2008]
- ► Non-negative matrix factorisation (NMF) [Lee & Seung, 2001]
- ► Gamma Poisson model (GaP) [Canny, 2004]

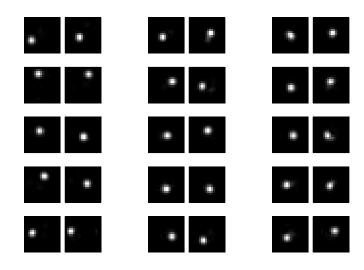
### Simulated Data

► Localised *W*(*x*)



### Simulated Data

▶ Separated W(x), N = 10



# Model comparison (estimation of K)

- 1. Principal component analysis (PCA)
- 2. Variational lower bound [Buntine & Jakulin, 2006]

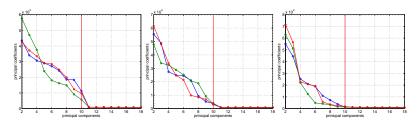
$$p(K|D) \propto p(D|K)p(K)$$

3. Analysis of the correlations in residuals

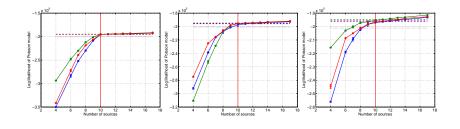
$$s_{xt} = \frac{d_{xt} - \sum_{k=1}^{K} w_{xk} h_{kt}}{\sqrt{\sum_{k=1}^{K} w_{xk} h_{kt}}}$$

# Model comparison (estimation of K)

► PCA

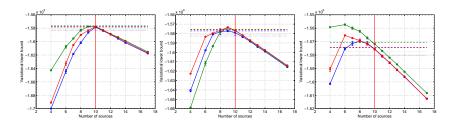


► Log likelihood

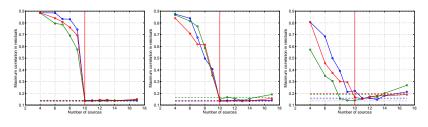


# Model comparison (estimation of K)

► Variational lower bound [Buntine & Jakulin, 2006]



Correlations in residuals (data - model)



### Theoretical limits for LM - Cramer Rao bound

lacktriangle Covariance matrix of an unbiased estimator heta

$$\mathsf{Q}(\theta) \geq \mathsf{I}^{-1}(\theta)$$

► Fisher information as a curvature of the log-likelihood function

$$\mathbf{I}_{ij}(\theta) = -\mathbb{E}\left[\frac{\partial^2 \mathcal{L}}{\partial \theta_i \partial \theta_j}\right]$$

#### Two sources

centres of the sources

$$\theta=(c_1,\,c_2)$$

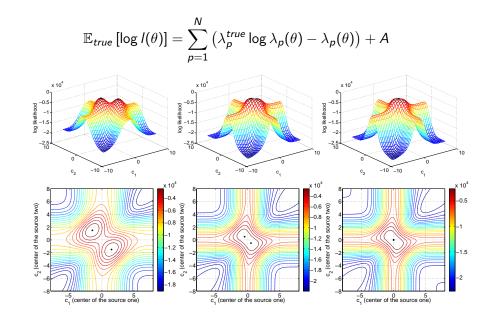
likelihood

$$I(\theta) = \prod_{p=1}^{N} Po(n_p | \lambda_p(\theta))$$

mean intensity image

$$\lambda(\theta) = \Lambda_1 f_1(c_1) + \Lambda_2 f_2(c_2) + b$$

## Fisher information as a curvature of the log-likelihood



## Blinking sources

blinking situation - intensity  $\Lambda = (\Lambda_1, \Lambda_2)$  is a random variable distributed over four states:

$$\left\{ {{\Lambda }^{1}}=\left( {{\Lambda }_{1}},0 \right),\,{{\Lambda }^{2}}=\left( {{0},{\Lambda }_{2}} \right),\,{{\Lambda }^{3}}=\left( {{\Lambda }_{1}},{{\Lambda }_{2}} \right),\,{{\Lambda }^{4}}=\left( {{0},0} \right) \right\}$$

likelihood

$$I(\theta) = \prod_{p=1}^{N} p(n_p|\theta) = \prod_{p=1}^{N} \sum_{i=1}^{4} p(n_p|\theta, \Lambda^i) p(\Lambda^i)$$

### Localisation precision - source intensity

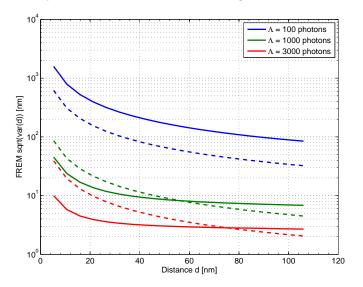


Figure: FREM (fixed background 100 photons)

## Localisation precision - source intensity

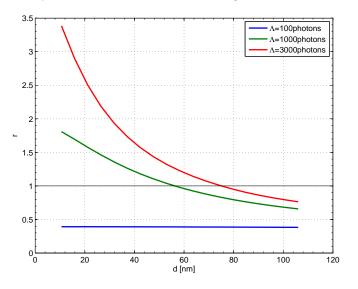


Figure: Comparison blinking vs static:  $r = \sqrt{\text{var}^{\text{static}}(d)/\text{var}(d)}$ 

## Localisation precision - background

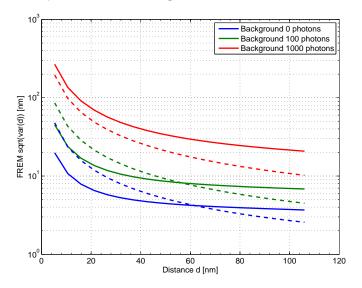


Figure: FREM (fixed  $\Lambda=10^3$  photons)

## Localisation precision - background

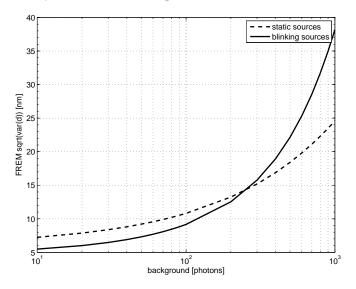
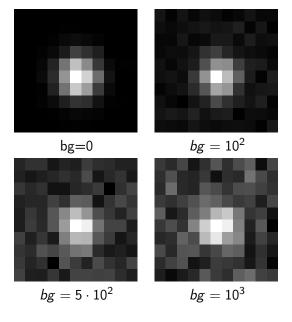


Figure: Effect of the background  $\Lambda = 10^3$  photons, d = 40 nm.

# Simulated data: Two points, $d=40 \text{ nm } \Lambda=10^3 \text{ photons}$



## "Resolution" of the correlation residual analysis

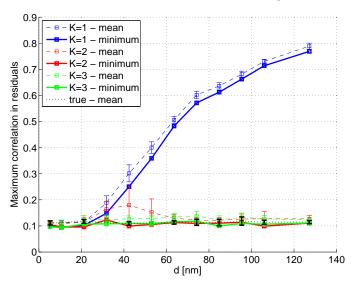
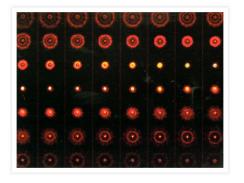


Figure:  $K_{true} = 2$ 

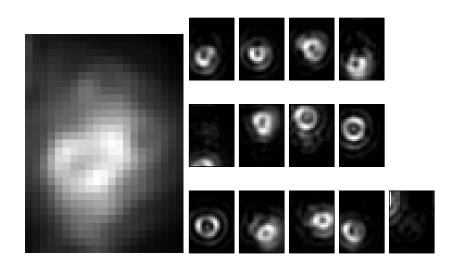
# Point Spread Function 3D



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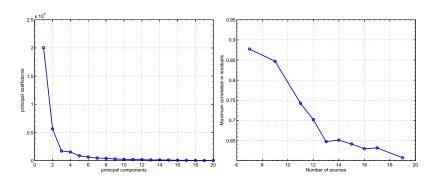
# Real data

▶ Separated W(x), N = 13



### Real data

▶ PCA vs Correlations in Residuals



### Future plans

- Write up: thesis chapters, paper (which journal?)
- ► More experimental work (visit of the Rainer's lab)
  - QD data, real sample
  - local illumination of the sample using spatial light modulator (SLM)
  - ► LM with photo switchable proteins