

A mixed-effects model approach to digital image analysis

M. Borelli Ph.D.

F.Zanconati, L.Bortolussi, F.Giudici, G.Barbati, L.Torelli
University of Trieste

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Statistics and histopathology

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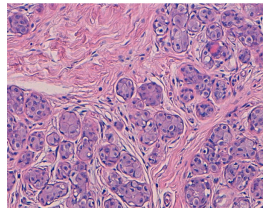
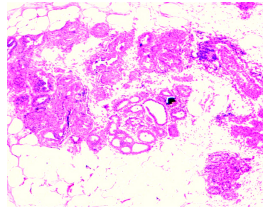
Histopathological images

Standard clinical practice

- cancer detection
- cancer classification

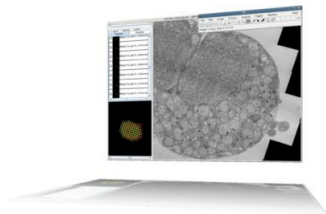
What to avoid

- inter-operator variability



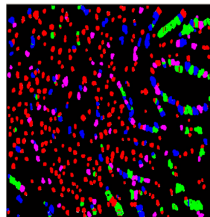
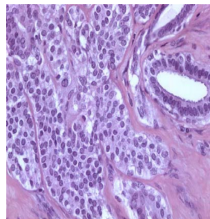
Manipulation softwares

- commercial products
 - ...
- open-source products
 - ...
 - Fiji
 - ...



Difficulties in *expert systems*

- low-level information
 - pixel values
- high-level information
 - relationships pixel/object
- domain-specific information
 - histo structures relationships



Milestones

- Dalle et al., IEEE-EMBS, 2008.
- Singh et al., IJEST 3(5), 2011.

Probabilistic / statistical approaches

- ...
- scoring methods
- bayesian classifiers
- neural networks
- ...

Our proposal

- mixed-effects models

What are mixed-effects models

Fixed effects + random effects

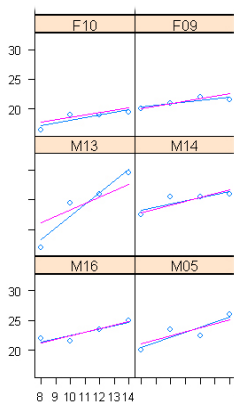
- repeated measures data
- longitudinal data
- hierarchical / clustered data

$$y_{ij} = \mu + b_i + \epsilon_{ij}$$

One of the advantages of mixed-effects models

The *shrinkage* phenomenon

- Strenio et al., Biometrics 39, 1983.

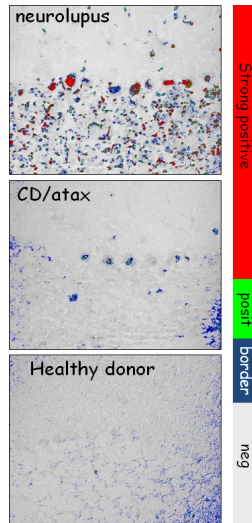


A previous result with mixed-effects modelling

Trieste Autoimmune Brain Atlas

- S. Zulian, M. Borelli, E. Tongiorgi. *Ongoing*

$$densito_{ij} = \beta_i + b_j + \epsilon_{ij}$$

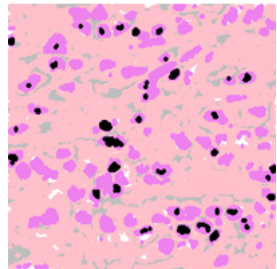
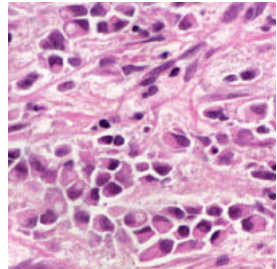


Open source softwares

- imagemagick
- R
 - library(pixmap)
 - library(lme4)
- bioconductor
 - graph

Results

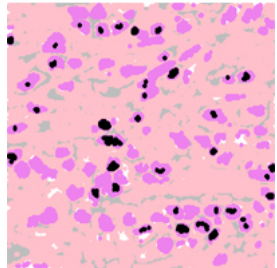
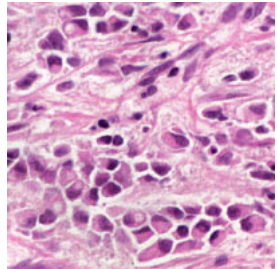
- detection of nuclei
 - Manhattan distance
- counts of nuclei
 - Depth first search algorithm



Cancer classification by mixed-effects modelling

Future steps

- supervised classification
 - to train a *learning machine*



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