



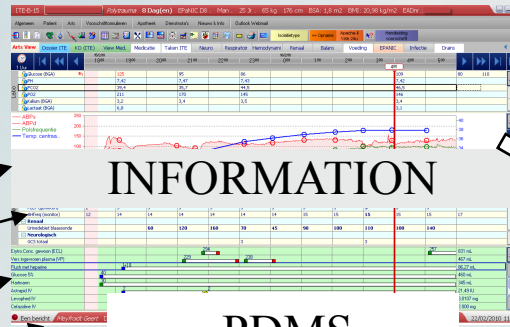
# Computerized data management in the ICU.

PREDICTIVE MODELING, TIME SERIES  
ANALYSIS AND OPPORTUNITIES FOR  
SUPPORT OF CARE.

Geert Meyfroidt  
SIZ award 2011

# Intensive care unit: data rich environment!

DATA:  
+/- 250 categories.



INFORMATION

PDMS

KNOWLEDGE:  
+/- 7 variables

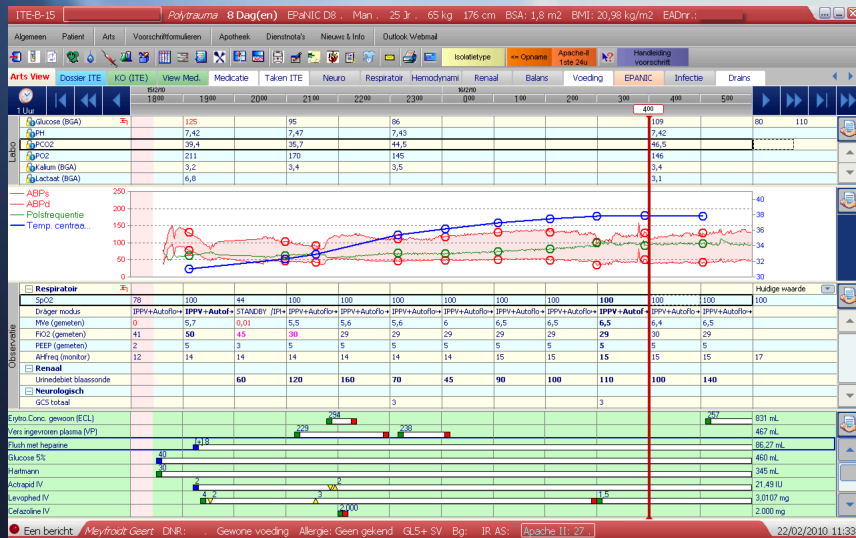


WISDOM

Ackoff RL: From Data to Wisdom. J Appl Syst Analis, 1989

Miller G: The magical number seven plus or minus two. Psychol review, 1956

# Patient data management system

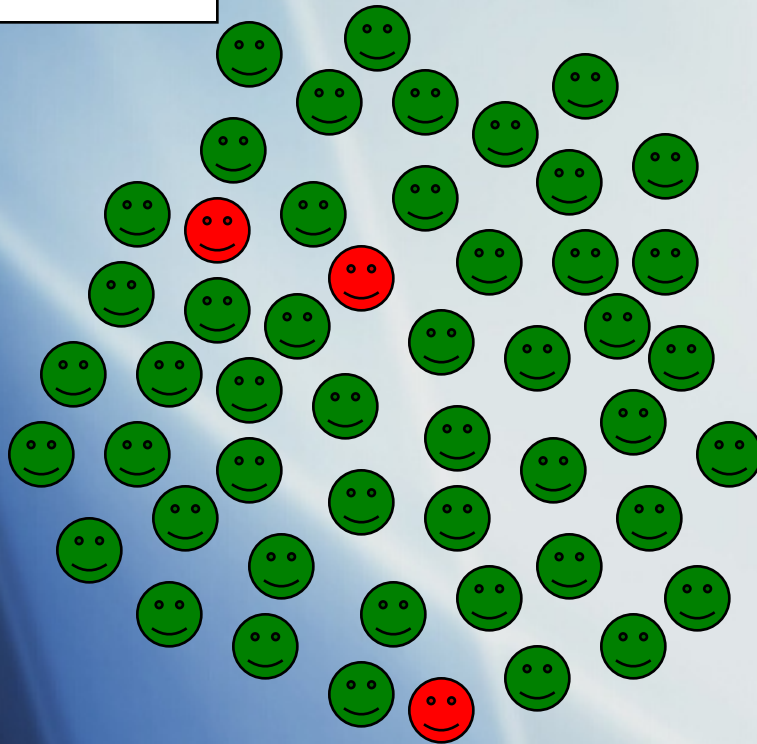


- “Reporting”
  - Monitors
  - Lab
  - Devices (MV,RRT, ...)
  - Observations
  - ....
- Prescription: CPOE
- Decision support



# Predictive modeling

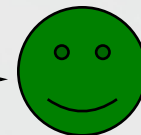
Observations



Outcome



Generalization  
"MODEL"



94%



"Learning"

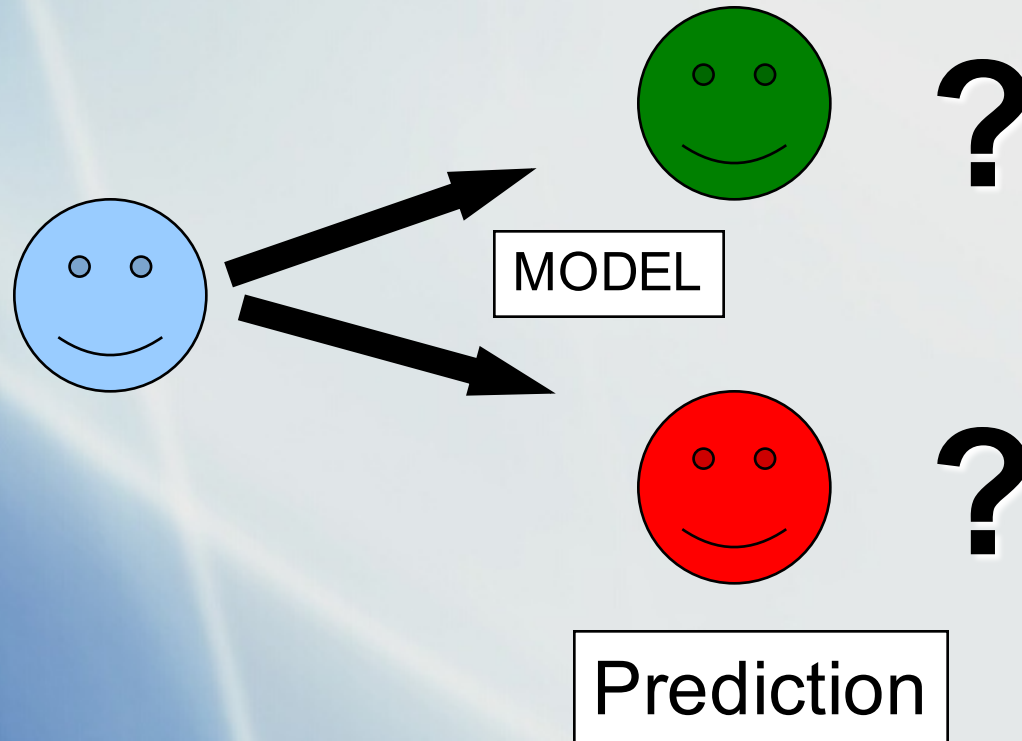


6%

Admission: demographics, medical history, admission diagnosis, ...  
Observations: lab, medication. ...  
Devices: monitor, ventilator, renal replacement therapy, ...

PDMS

# Predictive modeling



# Data mining and machine learning

- Automatic learning
- Incorporation of background knowledge
- Examples
  - Weather forecasts
  - Fraudulent bank transactions
  - IBM Watson project
  - Genome studies
  - ...

# Machine learning and ICU predictions: proof of concept



1548 patients  
-Admission data  
-"Once daily" data

## Predictive tasks:

1. ICU mortality
2. ICU LOS > 3 days
3. Development of renal failure
4. Recovery from renal failure

Meyfroidt G et al: *Best Pract & Res Clin Anaesth* 23 (2009) pp. 127-143

Ramon J, Fierens D, Güiza F, Meyfroidt G, et al. *Adv Eng Inf* (2007)21 (3) pp 243-256



# Machine learning and ICU predictions: proof of concept

aROC > 0.80

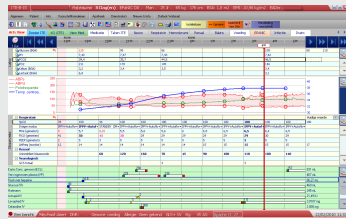
| Task                      | Day       | DT   | RF   | NB   | TAN  |
|---------------------------|-----------|------|------|------|------|
| Mortality                 | admission | 0.79 | 0.82 | 0.88 | 0.86 |
| ICU LOS > 3 d             | admission | 0.75 | 0.79 | 0.83 | 0.83 |
| Renal failure:development | n-2       | 0.82 | 0.86 | 0.86 | 0.88 |
|                           | n-3       | 0.84 | 0.87 | 0.87 | 0.88 |
|                           | n-4       | 0.83 | 0.88 | 0.87 | 0.87 |
| Renal failure:recovery    | n-1       | 0.82 | 0.87 | 0.80 | 0.85 |
|                           | n-2       | 0.80 | 0.84 | 0.78 | 0.84 |
|                           | n-3       | 0.80 | 0.85 | 0.80 | 0.87 |
|                           | n-4       | 0.81 | 0.86 | 0.81 | 0.87 |

(10-fold cross validation)

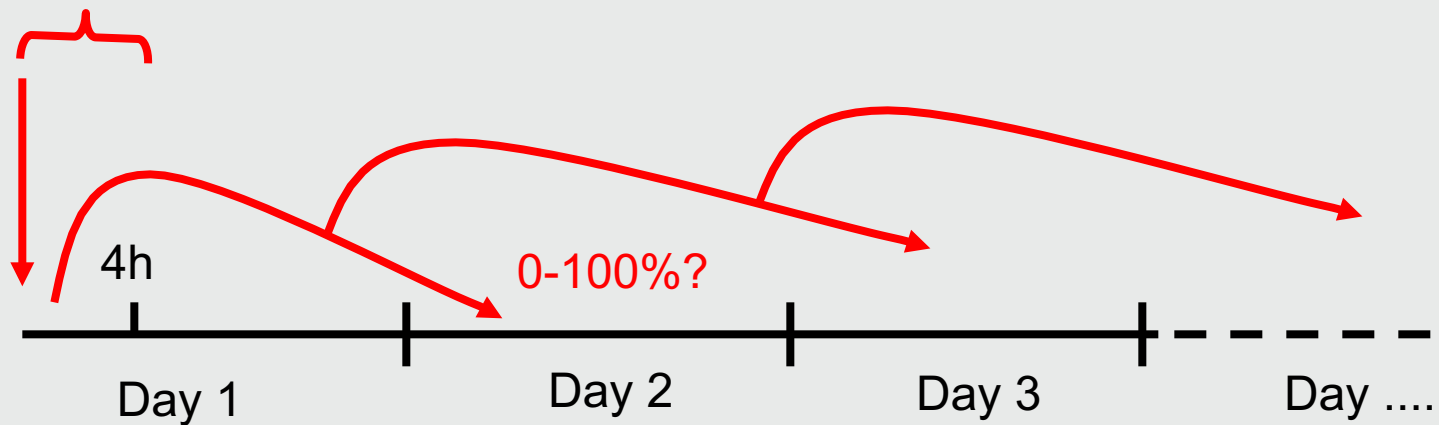
Meyfroidt G et al: *Best Pract & Res Clin Anaesth* 23 (2009) pp. 127-143

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# Predicting ICU length of stay after cardiac surgery



## 2. Prediction of the day of discharge



# Predicting ICU length of stay after cardiac surgery

Probability of discharge on the day after surgery?

|                            | aROC           | Brier Score    |
|----------------------------|----------------|----------------|
| Gaussian Processes (n=499) | >0.75          | <0.25          |
| EuroSCORE (n=499)          | 0.726 =        | 0.324 <b>S</b> |
| Nurse (6 u) (n=396)        | 0.695 <b>S</b> | 0.245 <b>S</b> |
| Physician (6 u) (n=159)    | 0.758 =        | 0.216 <b>S</b> |

$$BS = \text{Mean } [p_n - o_n]^2$$

# Predicting ICU length of stay after cardiac surgery

Prediction of the day of discharge?

|                            | LPF                      | LPF=0        | RMSRE |
|----------------------------|--------------------------|--------------|-------|
| Gaussian Processes (n=499) | 0 (0 - 0.4)              | 40%          | 0.408 |
| EuroSCORE (n=499)          | -0.3 (-0.5 - 0) <b>S</b> | 19% <b>S</b> | 0.643 |
| Nurse (6 u) (n=396)        | 0 (0 - 0.3) <b>=</b>     | 38% <b>S</b> | 0.522 |
| Physician (6 u) (n=159)    | 0.2 (0-0.4)              | 31%          | 0.612 |

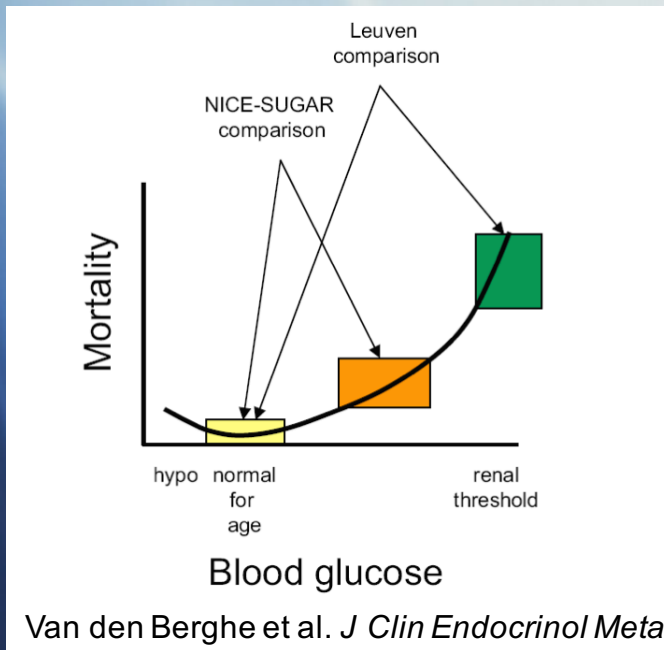
$$LPF = \frac{D_{\text{actual}} - D_{\text{predicted}}}{D_{\text{actual}}}$$

# Predicting ICU length of stay after cardiac surgery: conclusion

- Computer predicts better than Euroscore, and more reliably than clinician
- Models are not universally applicable but can be ‘relearned’ in a different clinical context.

# Tight glycemic control

Elevated blood sugar levels are associated with increased mortality

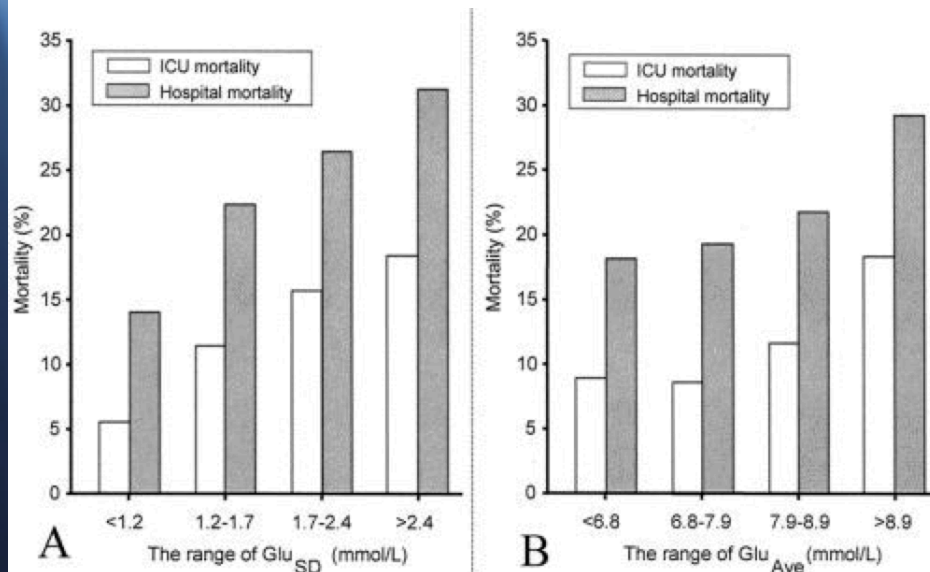


Van den Berghe et al, NEJM 2001  
Van den Berghe et al, NEJM 2006  
Vlasselaers et al, Lancet 2009  
NICE-SUGAR study, NEJM 2009

Normalization of glycemia with intensive insulin therapy reduces ICU mortality in certain clinical settings.

# Tight glycemc control

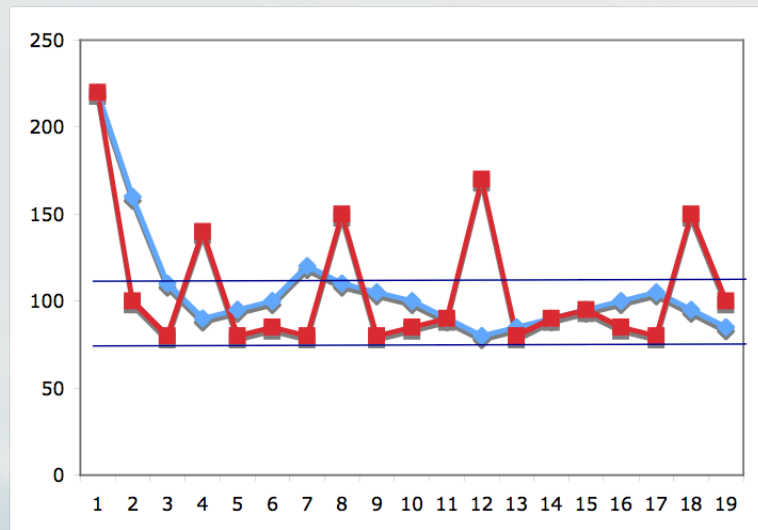
Increased blood glucose amplitude variability (BGAV) is associated with mortality



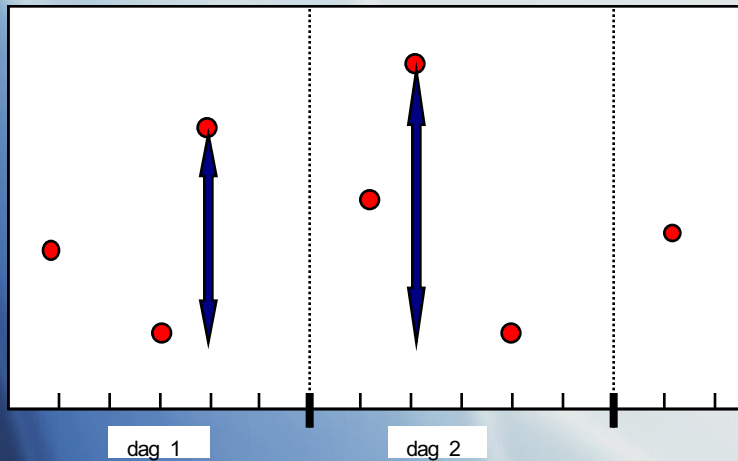
Egi et al, Anesthesiology 2006

Egi et al, Anesthesiology 2006  
Wintergerst et al, Pediatrics 2006  
Kransley et al, Crit Care Med 2008  
Dosset et al, Am J Surg 2008  
Ali et al, Crit Care Med 2008

Hyperglycemia-induced oxidative stress?



# Tight glycemic control: level and/or variability?



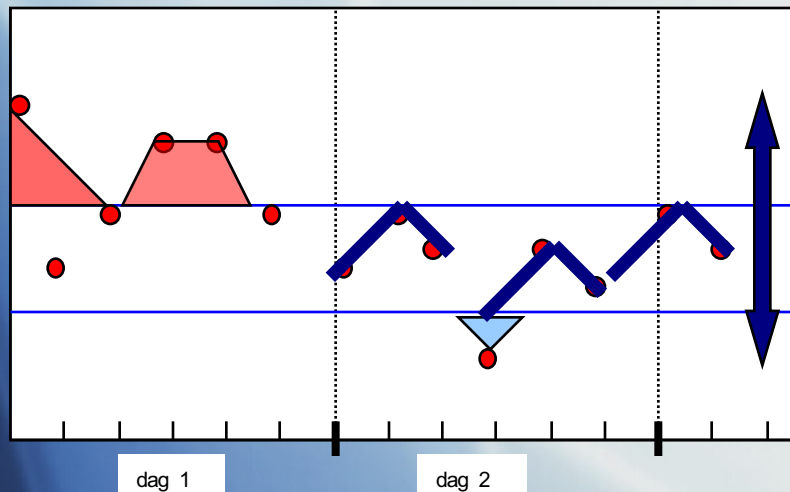
MICU+SICU: n=2748

**-level:** -mean morning  
-hypo

**-BGAV:** -mean daily  $\Delta$  BG



# Tight glycemic control: level and/or variability?



MICU: n=1200

|                  |        |
|------------------|--------|
| <b>-level:</b>   | -HGI   |
|                  | -HoGI  |
| <b>-BGAV:</b>    | -SD BG |
| <b>-pattern:</b> | -ApEn  |

# Tight glycemie control: level and/or variability?


|                 | Conventioneel    | Intensief         | P-value |                |
|-----------------|------------------|-------------------|---------|----------------|
| Mean morning BG | 151 (27)         | ↓ 104 (23)        | <0.0001 | } SICU<br>MICU |
| Hypoglycemie    | 2%               | ↑ 11%             | <0.0001 |                |
| Mean daily Δ BG | 59 (38-90)       | ↑ 72 (52-97)      | <0.0001 |                |
| HGI             | 58 (36-74)       | ↓ 14 (9-22)       | <0.0001 | } MICU         |
| HoGI            | 0.09 (0.00-0.36) | ↑ 0.90 (0.34-1,5) | <0.0001 |                |
| SD BG           | 38 (29-50)       | 36 (29-49) =      | 0.161   |                |
| ApEn            | 1.7 (0.3)        | 1.6 (0.3) =       | 0.126   |                |

Multivariabele  
log. regressie

MORTALITEIT

# Tight glycemetic control: impact of a computer alert

- IIT in Leuven
  - By nurses
  - Paper guideline (intranet)
    - Incomplete
    - Partially procedural/declarative
    - Informal



‘Intuitive decision making’  
No strict ‘if-then’ protocol

# Tight glycemetic control: impact of a computer alert

BG > 180 mg/dl  
Patient can eat

BG > 110 mg/dl  
IV- and/or enteral nu

BG 60-80 mg/dl







BG 40-60 mg/dl

BG < 40 mg/dl

Adapt insulin. New  
BG control in 1u

The screenshot shows a software interface for a medical alert. At the top, there are tabs for 'Event' and 'Details', and a status bar showing 'Waarde: 2' and 'Tijd: 04/09/2007 20:25'. The main alert area has a red header with the text 'Ev\_hypo40' and 'Kritiek!' (Critical), followed by a yellow progress bar. The alert message reads: 'De laatste glycemiewaarde is lager dan 40 mg/dl : Stop onmiddellijk de Actrapidrip , controleer de voeding en geef 10g Glucose IV . Controleer opnieuw na 30 min .' Below the message is a section titled 'Opmerkingen' (Remarks) with a list of items, each having a small icon and a dropdown arrow. A 'Bewaar' (Save) button is located at the bottom right of the interface.

# Tight glycemic control: impact of a computer alert

|                            | Pre-alert (729)<br>2/2007-7/2007 |   | Alert (644)<br>8/2007-2/2008 | P-value |
|----------------------------|----------------------------------|---|------------------------------|---------|
| <b>Blood glucose level</b> |                                  |   |                              |         |
| Mean BG                    | 112                              |    | 110                          | 0.002   |
| HGI                        | 10                               |    | 9                            | 0.004   |
| # BG values >110           | 33%                              |    | 30%                          | 0.008   |
| # BG values < 80           | 6%                               |   | 6%                           | 0.845   |
| # pat. with hypo < 40      | 6.5%                             |    | 4%                           | 0.043   |
| <b>BGAV</b>                |                                  |   |                              |         |
| SD BG                      | 28                               |  | 28                           | 0.566   |
| <b>BG monitoring</b>       |                                  |   |                              |         |
| # measurements/pt/d        | 7.7                              |  | 7.7                          | 0.891   |



# Computerized data management in the ICU

Conclusions

# Conclusions

- Advanced data analysis with automatically learning methods allows for the construction of customized clinically relevant predictive models.
- BGAV is an important feature that should be assessed when comparing blood glucose control strategies.
- Even a simple computer alert is able to have a significant impact on the quality of tight glycemic control.

# Present and future research

- ICU capacity planner for cardiac surgery
- Early warning alert for elevated ICP events in brain injured patients
- Prediction of bad outcome after brain injury
- Early warning alert for AKI
- Prediction of pharmacokinetics
- Time series summary statistics: morphological clustering
- Text mining: clinical notes



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