

### Selected Practical Aspects of Automated FHR Analysis

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### One slide about our lab



#### Czech Institute of Informatics, Robotics and Cybernetics



- Cyber-physical systems
- Intelligent systems
- Industrial informatics
- Robotics and machine perception
- Industrial production and automation
- Cognitive systems and neurosciences
- Biomedical and assistive technologies





CTG analysis group (<u>http://bio.felk.cvut.cz/ctg</u>)

- Václav Chudáček, Jiří Spilka automated CTG analysis, AI for FHR processing
- Michal Huptych, Miroslav Burša (FEE, CTU) data collection systems, clinical inf.
- Petr Janků, Lukáš Hruban (FN Brno) obstetricians at the University hospital in Brno





## **Context of the work**



### Intro: Current state of evaluation







FHR is measured by US or fECG

Signs of hypoxia are sought for

Decision are made based on FHR and clinical data

#### Outcomes:

- Healthy babies
- ✤ Caesarean sections (20-50% in CZ)
- Pathological pH (1-3%) -> severe cases may result in neurological damage such as cerebral palsy etc.



Machine representation of FHR is used





# Sources of motivation and frustration

"Wisdom comes from experience. Experience is often a result of lack of wisdom." - Terry Pratchett



### General



Automated analysis?

#### Ill defined inputs

- Databases in general small, and very difficult to compare
- CTG segment used for evaluation how long, how far, which stage
- ✤ Other inputs e.g. clinical, technical, etc.

#### Ill defined outputs

- What type of outcome do we use as an indicator of CTG/FHR class
  - pH+BDecf (reason: Cerebral palsy objective/statistical sound outcome measure): not very useful (very few cases)
  - Apgar5 (reason: ? current CPR techniques are able to get over 7 in 5th minute in most cases): unreliable (subjective, not necessary related to hypoxia)
  - pH alone (reason: it is easy): best (it does not relate to the long-term outcome, about 20% of measurements are wrong, no clear threshold)
  - Expert evaluation (reason: that is how it is done in the hospital): -(large variability, increased defensiveness in evaluation)

### **Clinical**



#### Hard to assess real-life CTG evaluation

- Interventions due to multiple factors
- CS with normal objective outcome (how often was it necessary?)
- Low-risk pregnancies not monitored
- ✤ Bad outcomes often related to very low quality or missing recordings

#### Guidelines in intrapartum CTG not very good regarding:

- Clear explanation of reasoning
- Consistency, Repeatability
- Reasons for use (cf. history of the CTG introduction)

Increase of CS without impact on # of hypoxic babies

Existence of many related confounding factors

Variability among fetuses (e.g. male vs. female)



### **Technical**



#### Majority of works done on static evaluation only

- One window (usually length < 40min, sleep episodes not considered)</p>
- Often no prediction (evaluation of full data)

#### Limits of technical methods not advertised

- Ultrasound or STAN recordings huge difference in terms of quality
- Some methods need certain length of recording without noise
- Tocographic data hardly usable

#### Lack of pathological cases

Preprocessing of the data is feature-dependent

Almost none of the other information available in the hospital is not considered



### **Pieces of our work**

#### (addressing in part the general problems above)





### **Database** CTU-UHB cardiotocograpic database



### **CTU-UHB database**



Collaboration with Ob&Gyn clinic in Brno

USG and STAN data

#### Only mature fetuses

First open-access CTG database

Common ground for algorithm comparison

Available at Physionet



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Chudáček et al. Open access intrapartum CTG database, BMC Pregnancy and Childbirth, 2014



#### Subjective – expert evaluation

- Annotations acquired via CTG Annotator
- Majority voting, Latent class model based on 9 experts
- Apgar score

Objective
 pH, pCO<sub>2</sub>, BDecf, BE

Mixture

✤ Majority, LCMs





## **FHR Signal processing**



### Signal pre-processing

#### Gap & Artefact detection

- ✤ Gap removal (< 15s)</p>
- Artefact rejection
- Bernardes inspired thresholds
- ✤ Adapted to 4Hz from beat to beat





Official obstetrics guidelines for CTG evaluation
 Circular definition of Acceleration/Deceleration
 Baseline detection based on histogram assessment







#### Morphological features (FIGO) (5)

Time-domain (6)
Freq.-domain (13)
HRV (4)
Wavelet (15)
Nonlinear (12)

In total 55 features



scale gives value of R for each variable pair





### Classification



### **Classification**





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### **Classification (3)**

#### Selected features

- Low spectral bands
- Decelerations
- Poincare plot SD2





### **Results – obj. annotation (2)**



### **Comparison of results**







♦ We need a common ground to compare

- Getting the data is cumbersome and unrewarding job
- But most hospitals now have electronic CTG records
- There is no "perfect database" yet

Experience shows that joining different approaches brings improvement across the board (kaggle.com)

Individual phase followed by joint effort phase

Could we build one together?



### **Outcome measures**

#### **Man-machine comparison**



### **Clinical evaluation vs. pH**

- Real-life CTG evaluation comparison vs. pH
- Variability
- Overall poor sensitivity
- Sensitivity drops Step3 -> Step4

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|-----|----|-----|---|
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| Annotation          | Objective | Step 1     |            | Step 2     |            | Step 3      |            | Step 4     |            |
|---------------------|-----------|------------|------------|------------|------------|-------------|------------|------------|------------|
|                     |           | SE         | SP         | SE         | SP         | SE          | SP         | SE         | SP         |
| majority<br>voting  | pH ≤ 7.05 | 29 (12-54) | 92 (88-95) | 41 (20-65) | 86 (81-90) | 86 (45-99)  | 86 (79-90) | 38 (18-63) | 94 (91-97) |
|                     | BD ≥ 12   | 30 (9-62)  | 92 (87-95) | 50 (22-78) | 86 (81-90) | 50 (3-94)   | 83 (76-88) | 22 (4-56)  | 93 (89-96) |
|                     | Apgar < 7 | 50 (10-90) | 91 (87-94) | 50 (10-90) | 85 (80-89) | 100 (5-100) | 83 (76-89) | 75 (25-99) | 93 (90-96) |
| hospital<br>records | pH ≤ 7.05 | 41 (20-65) | 94 (91-97) | 41 (20-65) | 94 (90-97) | 40 (15-71)  | 93 (88-96) | N/A        | N/A        |
|                     | BD ≥ 12   | 60 (29-85) | 94 (91-97) | 60 (29-85) | 94 (90-96) | 25 (1-75)   | 92 (87-95) | N/A        | N/A        |
|                     | Apgar < 7 | 0 (0-53)   | 92 (88-95) | 0 (0-53)   | 92 (87-95) | 33 (2-86)   | 92 (87-95) | N/A        | N/A        |

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Hruban et al. Agreement on CTG intrapartum recordings between expert-obstetricians, J. of Eval. in Clinical Practice, 2015



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Why?





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The latent class analysis (LCA) is used to estimate the true (unknown/hidden) evaluation of CTG and to infer weights of individual clinicians' evaluation – the latent class model (LCM).

The LCM considers clinical evaluation as a finite mixture of multinomial distributions.

Finite mixture models have fixed number of parameters and the standard method to estimate these parameters is expectation maximization (EM) algorithm.



### Latent class models









Biomedical Data Processing G r o u p

Spilka et al. Analysis of obstetricians' decision making on CTG recordings, J. of Biomed. Informatics, 2014



## Case studies – different experiments / projects in the field of CTG processing





### "Case study " 1: Scattering transformation

### (with Patrice Abry@ENSL, Stephane Mallat@ENS)



### **Scattering transform**

Introduced by S. Mallat
(http://www.di.ens.fr/data/scattering/)



### Scattering transform (2)



#### Wavelet transform



**\diamond** Complex mother wavelet  $\psi(t)$ 

♦ Dilated and translated wavelets  $\psi_{j,k}(t) = 2^{-j}\psi(2^{-j}(t-k))$ 

• Wavelet coefficients  $X \star \psi_{j,k}$ 

First-order coefs: Local time averages of abs. value of wavelet coefs.

$$SX(j,k) = \mathbb{E}\left\{ |X \star \psi_j| \right\} \approx N^{-1} \sum_{l=k}^{k+N} |X \star \psi_{j,l}|$$





### **Scattering Transform (3)**

Second order -> beyond Wavelets

 $\hat{\phi}_{J}(\omega) \uparrow |X \star \psi_{j_1}|(\omega) \quad \hat{\psi}_{j_2}(\omega)$ 

/avelet transform of absolute values of way

♦ Wavelet transform of absolute values of wavelet coefs.  $SX(j_1, j_2) = \mathbb{E} \{ ||X \star \psi_{j_1}| \star \psi_{j_2}| \} \approx N^{-1} \sum_{t=1}^{N} ||X \star \psi_{j_1}| \star \psi_{j_2}(t)|$ 

2<sup>nd</sup> order renormalized by the first

$$\widetilde{S}X(j_1, j_2) = \frac{SX(j_1, j_2)}{SX(j_1)} \approx \frac{\sum_{t=1}^{2^J} ||X \star \psi_{j_1}| \star \psi_{j_2}(t)|}{\sum_{t=1}^{2^J} |X \star \psi_{j_1}(t)|}$$

Nonlinear transform:

- ✤ Goes beyond wavelet
- Explores dependencies beyond correlation (or spectrum)



### **Relation scattering - scaling**



\*Relation between scattering and scaling  $SX(j,k) \sim 2^{jH}$ 

- H Hurst exponent
- $z(j_1)$  scaling exponents that may depart from H

$$\widetilde{S}X(j_1,j_2) \sim 2^{(j_1-j_2)z(j_1)}$$


# **Fractal Dynamics of FHR**



#### First order



### Fractal behaviour:

♦ Time scales ranging from  $4s < a = 2^j < 60s$ 



# **Fractal Dynamics of FHR (2)**

Second order for  $j_1 = 1, 2, 3$ .



### Fractal behaviour:

♦ Time scales ranging from  $4s < a = 2^j < 60s$ 

## **Results**



#### Discrimination power on SDB (HFME Lyon)



#### Performance outcome





# "Case study " 2: Scaling properties of FHR

## (with Patrice Abry@ENS)



# **Temporal dynamics**



### Classical measures

- ✤ STV scale of a = 3.75s (antepartum)
- ✤ LTV scale of a = 60s (intrapartum)

## Why to limit ourselves to these arbitrary intervals?



# **Properties of FHR**





## "Spectrum" : $\Gamma_X(f) \sim C|f|^{-(2H-1)}, |f| \rightarrow 0$



# **Continuous Wavelet Transform**

$$X(t) \to T_x(a, t) = \langle \frac{1}{a} \psi \left( \frac{u-t}{a} \right) | X \rangle$$



Joint time and frequency energy content



# **Fractal exponents**



- Oscilations -> wavelet coefs.
- Variability is not characterized by actual value
- Scale invariance is measured instead -> H
- The H computed via wavelet spectrum provides
  - Variability at all scales jointly (not just STV/LTV scale)
  - ✤ Gives information of temporal dynamic of HF/LF ratio

# **Scale invariance in FHR**





Power Law Behavior:  $a_m = 2^3 \le a \le a_M = 2^7$ 

# **Values of H per class**





# **Influence of decelerations**





# **Influence of decels. on H**





# Conclusion



### Hurst exponent

- Allows representation of time-scale properties of FHR by a single value
- Measures embraces the Temporal Dynamics as Fractal Variability
- ✤ Gathers time and spectral variabilities of the FHR in one feature
- Describes temporal dynamics across range of scales rather than for specific scales
- ✤ Simplifies the FHR analysis (in contrast to e.g. FIGO)

Behaves consistently irrespective to decelerations





# "Case study " 3: Mobile CTG

## (with Siemens A.G. Austria)



# **General schema of mCTG**





# **Phonography signal processing**







# "Case study " 4: Latent class model

## (by Jiří Spilka@CTU)



# **Motivation**





# **Results on pH**





- used 3 class scenario
- black squares false negatives
- pathological  $pH \le 7.10$
- suspicious pH (7.10, 7.15]
- normal: pH > 7.15
- misclassification near boundary
- #16 recs. not pathological BDecf
- #11 recs. high Apgar score  $\geq$  9 (max. value 10)
- #11 recs. for each record  $\geq$  5 clinicians said FHR is normal



## Latent class model (LCM)



Input data: noisy and imprecise

• 
$$y_i^j$$
,  $j = 1, ..., J$ ,  $i = 1, ..., N$ 

• Goal: estimate ground (unobserved) truth

$$\Pr[\mathbf{y}_i^1, \dots, \mathbf{y}_i^J | \boldsymbol{\theta}] = \prod_{i=1}^N \left[ \sum_{c=1}^C \pi_c \Pr[\mathbf{y}_i^1, \dots, \mathbf{y}_i^J | \boldsymbol{\theta}_c] \right]$$





# **Results on pH with LCM**

## LCM – false negatives from supervised learning



- no strict boundary: normal ∘, suspicious △, pathological □
- false negatives (

   score, clinical evaluation



# "Case study " 5: OB information system – The Delivery Book

(by Michal Huptych@CTU)



## **Electronic delivery book**



#### The application has been deployed as pilot version.

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| Presidence     Nasiedugel #       a)       Jimeno rodicky: Ggg Ggg, 6. porodopisu: 111232       dicka i Příjeni     Porod       Diegnózy/Operace/Pr       OrožetneC       Cislo nover.     502       Polsa       Défaa       11.50       Pasie       Défaa       11.51       Défaa       11.54       Défaa       11.54  | ecounici Noverezemec   | Nesrutologie  | Blos<br>Jméno rodičky: Ggg<br>Rodička / Přijen<br>Neonatologie<br>1. Detam přijeti<br>2. Prozentovate<br>2. Prozentovate<br>1. Detam přijeti<br>1. De | Gigg, 6. portodopiau: 11<br>Portod D<br>06.02.2014                             | b)<br>1233<br>angriedzy/Operace/f                      | Processici<br>Diagnotic pri<br>propulsion<br>Propulsion<br>des million res<br>Dispontantance | Novorozenec<br>atrosresta ener  | Neonatologie<br>+<br>trost.cov mosta 0050<br>+<br>trost.cov mosta 0050<br>+<br>-<br>-<br>-<br>-<br>-<br>-<br>-<br>-<br>-<br>-<br>-<br>-<br>- |
| Predichop     Nasiedugici i       Jindeno rodicky: Gog Gog, 6. porodopisu: 111202       dicka ( F/jern     Pored       Didenocyc/Gperace/Pr       Obligenocyc/Gperace/Pr  | RECOVINES Novereasement  | Nesnustoiogie | Akor<br>Jméno rodičky: Ggg<br>Rodička i Prijem<br>Neonatologie<br>1. Datam prijeb<br>seconecost<br>2. Treamatismus<br>Hespune JM<br>Hitubase<br>Hrose<br>Hitubase   | Ggg, č. porodopisu: 11<br>Porod D<br>06.02.0014                                | b)<br>1233<br>augriozy/Operace/F                       | Pracovski<br>Diagnolog při<br>propublikel<br>Propublikel<br>Diagno-curicace                  | Novorozenec<br>Aly mattormace pl<br>Atronmatika cher<br>+<br>-<br>-<br>Mantingie<br>Meanologie  | Neonatologie<br>thras.cav mosku 0000<br>mosomik, H 5 Optis<br>194 24 Solentiel   |
| Presidence     Nestedușci i       Jimeno rodicky: Geg Geg, c. porodopisu: 111202       oddără i Prijeri     Pored       Oblizia     Oblizia       Oblizia     07       Oblizia     07       Vero zender     288       Oblizia     07       Vero zender     288       Apgar     0       Internation     7.21       Minse 0-18     7.34       Minse 0-19     7.34   | ecovnici Noverezenec   | Necrustologie | Stor<br>Jmeno roditky: Ggg<br>Roditka (Pfijen<br>Neonatologie<br>1. Detam physi<br>meaning  | Gigg, 6. porediopless: 11<br>Pered D<br>06.02.2014                             | b)<br>1223<br>augriczy/Operacoff<br>a normał<br>-<br>- | Precovnici<br>Diagoniza při<br>propublikní<br>Propublikní<br>Diapontanizace                  | Novorozenec<br>Al martineze p<br>Almartineze p<br>e<br>e<br>os.oz.zo<br>Plantineze<br>Martineze | Neonatologie   |
| Presidences     Nasiedagel I       Jimeno rodicky: Gog Gog, c. porodopisu: 111232       picka i Příjeni     Porod       Dicka i Příjeni     Stana •       Dicka i Příjeni     222       Apgar     0       Bit artaritini     7,21       Bit artaritini     1,14       Bit artaritini     1,24       Příjenení dikíte     0  | ecounici Noverezenec   | Nesrustologie | Alear<br>Jméno rodičky: Ggg<br>Rodička / Přijen<br>Neonatologie<br>1. Dotan přijeti<br>2. Prozestana<br>1. Dotan přijeti<br>1. Do | Gigg, C. portedoplas:: 11<br>Perced D<br>06.02.0014<br>Doba 9v2ni<br>risateirn | b)<br>1253<br>segricEp/Operace/F<br>all normal<br>ta   | Processici<br>Diagnolog při<br>propuštění<br>Propuštění<br>do měcí na<br>Diapentaritace      | Noverozenec<br>Alv mathomace of<br>Altromatika ether  | Neonatologie   |



## Domain model in obstetrics







# "Case study " 6: DeBo database mining Epidemiological study of pH

## (with Ibrahim Abou Khashbah@CIIRC)





### 12 878 patients from FN Brno

## **◆**99 pH < 7.05, 231 pH <7.1

### 107 features

| Feature                           | n(-) | $\operatorname{prct}(\%)$ | OR (95% CI)              |
|-----------------------------------|------|---------------------------|--------------------------|
| <b>pH</b> ( $\leq 7.1$ )          | 251  | 2.36                      | -                        |
| Apgar score 5min $(< 8)$          | 203  | 1.91                      | $12.08 \ (8.33 - 17.53)$ |
| Sectio Caesarea                   | 2110 | 20.30                     | $1.12 \ (0.83 - 1.52)$   |
| Induced delivery                  | 2198 | 20.69                     | 1.82(1.39 - 2.39)        |
| Entonox(medic.)                   | 144  | 1.36                      | $1.18 \ (0.43 - 3.22)$   |
| Epidural analg.                   | 1915 | 18.03                     | $1.25\ (0.92 - 1.70)$    |
| Ist stage ( $> 360$ min)          | 1807 | 18.40                     | $1.40 \ (1.04 - 1.89)$   |
| IInd stage ( $> 30$ min)          | 884  | 10.11                     | 2.87 (2.07 - 3.97)       |
| Parity $(< 2)$                    | 5299 | 49.99                     | $3.42 \ (2.54 - 4.59)$   |
| Sex (Male)                        | 5468 | 51.53                     | $0.83 \ (0.65 - 1.07)$   |
| O100 – hypertension               | 533  | 5.02                      | $1.30\ (0.78 - 2.17)$    |
| O140 – preeclampsia               | 114  | 1.07                      | $0.74\ (0.18 - 3.00)$    |
| O365 - IUGR                       | 375  | 3.53                      | $1.14 \ (0.60 - 2.16)$   |
| O681 – meconium                   | 784  | 7.38                      | $1.60 \ (1.07 - 2.38)$   |
| D650 – defibrination syndrome     | 258  | 2.43                      | $1.33 \ (0.65 - 2.73)$   |
| D695 – secondary thrombocytopenia | 36   | 0.34                      | 3.79(1.15 - 12.44)       |



# "Case study " 7: CTG Trainer App for Android and web

## (with Petr Stuchlik@PPF)









Naměřené hodnoty

| Ddhlásit |  | Dostupný test | Má CTG signál    |     | nál      | Příjmení |       |                                 | _              |            |
|----------|--|---------------|------------------|-----|----------|----------|-------|---------------------------------|----------------|------------|
|          |  | - nezáleží -  | •                | ano | •        |          |       |                                 | Reset          |            |
|          |  |               | Název            | G   | ravidita | Parita   | Týden | Datum a čas porodu <del>v</del> | IČ chorobopisu | Pat. porod |
|          |  | O Prohlížet   | Kamila Vašourko  | ová | IV       |          | 40    | 06.10.2014 - 14:00              | 1319003        | ne         |
|          |  | O Prohlížet   | lveta Švančarová | 5   | V        | N        | 41    | 06.10.2014 - 13:25              | 1318856        | ne         |
|          |  | O Prohližet   | Martina Pospíšik | ová | IV       |          | 38    | 06.10.2014 - 13:15              | 1318828        | ne         |
|          |  | O Prohlížet   | Jana Vachalová   |     |          |          | 36    | 06.10.2014 - 12:45              | 1318823        | ne         |
|          |  | O Prohlížet   | Denisa Bartoñov  | rá  | 1        | 1        | 40    | 06.10.2014 - 12:40              | 1318961        | ne         |
|          |  | O Prohlížet   | Petra Rybnikářov | vá  | ш        |          | 40    | 06.10.2014 - 12:30              | 1318834        | ne         |
|          |  | O Prohlížet   | Dana Doleželová  |     | 1        | 1        | 41    | 06.10.2014 - 12:30              | 1318829        | ne         |
|          |  | O Prohlížet   | Darina Magdová   |     | ш        |          | 37    | 06.10.2014 - 10:50              | 1316169        | ano        |
|          |  | O Prohližet   | Kateřina Slováko | ová |          |          | 40    | 06.10.2014 - 10:45              | 1318808        | ne         |
|          |  | O Prohlížet   | Monika Štodrová  | é . |          |          | 39    | 06.10.2014 - 09:55              | 1318830        | ne         |
|          |  | O Prohlížet   | Monika Pelouško  | ová | 1        | 1        | 38    | 06.10.2014 - 09:00              | 1318825        | ne         |
|          |  | O Prohlížet   | Petra Slavíková  |     | 1        | 1        | 37    | 06.10.2014 - 05:20              | 1318817        | ne         |
|          |  | O Prohlížet   | Valeria Matyáš   |     | 1        | 1        | 39    | 06.10.2014 - 04:55              | 1318718        | ne         |
|          |  | O Prohližet   | Veronika Jinová  |     |          |          | 39    | 06.10.2014 - 04:35              | 1318820        | ne         |
|          |  | O Prohlížet   | Markéta Matušk   | ová |          |          | 40    | 05.10.2014 - 18:30              | 1318791        | ne         |
|          |  | O Prohlížet   | Ivana Horňáková  | ò   |          | 1        | 40    | 05.10.2014 - 18:00              | 1318772        | ne         |
|          |  | O Prohližet   | lveta Řičánková  |     | 1        | 1        | 39    | 05.10.2014 - 17:15              | 1318734        | ne         |
|          |  | O Prohlížet   | Hanna Kryshtof   |     | 1        | 1        | 40    | 05.10.2014 - 17:10              | 1318730        | ne         |
|          |  | O Prohlížet   | Lenka Havličkovi | ó   | 1        |          | 40    | 05.10.2014 - 16:00              | 1318487        | ne         |
|          |  | O Prohlížet   | Martina Houdko   | rvá |          | 1        | 38    | 05.10.2014 - 14:05              | 1318688        | ne         |



#### 🚯 KTG+ Fyziologie - Interpretace KTG - Prohlížeč KTG Analýza STAN - Databáze záznamů Doporučení ČGPS

#### KTG+ Viewer

#### CTG Viewer demo I. (1308098)

Věk: 32, GRAV: II, PARA: I, týden: 32 + 1, indukovaný porod: ne







#### **KTG+ Viewer**



р







2

9/6/2014

| Interno pacienta               | Vilk matky | Gravidita | Parita | Týden tilhotenatul | pH   | C Test |
|--------------------------------|------------|-----------|--------|--------------------|------|--------|
| udela Vodova (1040209)         |            | 1         | 1      | 40                 | -1.0 | NE     |
| teamika Mishea (1279426)       | 33         |           | 1      | 40                 | 7.23 | NE     |
| terbora Borancová (1043120)    |            |           | 1      | 38                 | 7.87 | NE     |
| Banka Vacková (1107343)        |            | 1         | 1      | 38                 | 7.25 | NE     |
| Dana Skobodová (1068597)       |            | 1         | - C    | 40                 | 7.23 | NE     |
| Niz Stałkowa (1142666)         |            |           | i.     | 39                 | 72   | NE     |
| Doubravka Rožnovská (1299723)  | 34         | 1         | 1      | 40                 | 7.14 | NE     |
| liken Kuchyfikosá (1178254)    |            | w         | н      | 41                 | 7.19 | NE     |
| iva Marková (1289280)          | 35         |           | 1      | -40                | 7.02 | NE     |
| labriela Petificová (1307653)  | 34         | 1         | E.     | 38                 | 7.3  | NE     |
| kana Benková (1023687)         |            |           | 1      | 40                 | 7.47 | NE     |
| amila Hyzikiová (1141563)      |            |           | 1      | 29                 | 7.13 | NE     |
| Itia Bočková (1072414)         |            | 1         | 1      | 39                 | 7.14 | ANO    |
| Canelina Dokoupilová (1249990) |            |           | 1      | 39                 | 7.14 | AND    |
| Laterlina Politiková (1284077) | 24         | 1         | 1      | 40                 | -1.0 | NE     |
| Cristýma Rejtharová (1294463)  | 25         | 1         | ÷.     | 40                 | 7.17 | AND    |



# "Case study " 8: Clustering of FHR using SAX

## (with Helen Drosou@TEI of Athens)





#### Can we get by without those anoying features?

SAX – symbolic approximation

- Based on piecewise agregate approximation
- Simple k-means algorithm is used for clustering
- Preliminary results very promising given there are no "classical" features involved





# "Case study " 9: Embedding & graph of graphs

## (with Patrice Abry@ENS, Ronen Talmon@Technion)








### "Case study " 10: New biochemical markers

#### (with Laboratory of Molecular Diagnostics FN Brno)





#### New markers

#### ✤ mRNA

 Messenger RNA (mRNA) is a large family of <u>RNA</u> molecules that convey <u>genetic information</u> from <u>DNA</u> to the <u>ribosome</u>, where they specify the <u>amino acid</u> sequence of the <u>protein</u> products of <u>gene</u> <u>expression</u>.

#### ✤ miRNA

- A microRNA (abbreviated miRNA) is a small non-coding RNA molecule (containing about 22 nucleotides) found in plants, animals, and some viruses, which functions in RNA silencing and post-transcriptional regulation of gene expression
- The <u>human genome</u> may encode over 1000 miRNAs, which are abundant in many mammalian cell types and appear to target about 60% of the genes of humans and other mammals.







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- Lukáš Zach, Petr Stuchlík BSc./MSc. students, programmers

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- Lukáš Hruban, Martin Huser
  - Part of the IGA team in FH Brno
- Michal Koucký
  - Obstetrician, 1st Medical Faculty Charles University in Prague.
  - First contact with the topic (2008)











# Thank you for your attention!









# Variability & HF/LF features united Scaling properties of FHR



#### **Temporal dynamics**



#### Classical measures

- ✤ STV scale of a = 3.75s (antepartum)
- ✤ LTV scale of a = 60s (intrapartum)

#### Why to limit ourselves to these arbitrary intervals?





### **Continuous Wavelet Transform**

$$X(t) \to T_x(a, t) = \langle \frac{1}{a} \psi \left( \frac{u-t}{a} \right) | X \rangle$$



Joint time and frequency energy content



#### **Fractal exponents**



- Oscilations -> wavelet coefs.
- Variability is not characterized by actual value
- Scale invariance is measured instead -> H
- The H computed via wavelet spectrum provides
  - Variability at all scales jointly (not just STV/LTV scale)
  - ✤ Gives information of temporal dynamic of HF/LF ratio

### **Scale invariance in FHR**





Power Law Behavior:  $a_m = 2^3 \le a \le a_M = 2^7$ 

#### **Values of H per class**





### **Influence of decelerations**





### **Influence of decels. on H**





### Conclusion



#### Hurst exponent

- Allows representation of time-scale properties of FHR by a single value
- Measures embraces the Temporal Dynamics as Fractal Variability
- ✤ Gathers time and spectral variabilities of the FHR in one feature
- Describes temporal dynamics across range of scales rather than for specific scales
- ✤ Simplifies the FHR analysis (in contrast to e.g. FIGO)

Behaves consistently irrespective to decelerations







- HF and LF bands set-up taken from adults
- No evidence proved the threshold to be universal
- Could we use other thresholds?
- Is the band splitting relevant, then?
- H can be robust alternative to HF/LF

| $f_{interm}$ | AUC         | p-value |
|--------------|-------------|---------|
| 0.10         | 0.79 (0.08) | < 0.05  |
| 0.15         | 0.82 (0.07) | < 0.05  |
| 0.20         | 0.82 (0.07) | < 0.05  |
| 0.25         | 0.80 (0.07) | < 0.05  |
| 0.30         | 0.77 (0.08) | < 0.05  |
| 0.35         | 0.78 (0.08) | < 0.05  |







### **Power law behaviour on classes**



ta Processing G r o u p

### What is the goal?

#### CTG-centric: Related mainly to CTG

- Correctly classified CTG (based on guidelines)
- Detection of abnormal CTG => outlier detection
- Classification into classes based on data => clustering

#### Baby-centric: Related to delivery as a whole

- Always healthy baby => rate of CS close to 100%
- "Happy" baby & mother
- Good pH (or any other "objective") value

#### M.D. centric:

- Decision based on objective rather than subjective indices
- Less legal trouble for obstetricians









### **Problems with goals (1)**



#### CTG-centric: Related mainly to CTG

- ✤ Relation of CTG and outcome (Se is ~40%, Hruban2014 submitted)
- Correctly classified CTG (based on guidelines)
  - Guidelines are flawed and irreproducible (Campos2010)
  - Guidelines are taking over common sense (Ugwumadu2014)
- Detection of abnormal CTG => outlier detection
  - In theory sound idea yet about 30-50% of pathological cases misclassified (unpublished results on HFMEdb of about 3000 cases)
- Classification into classes based on data => clustering
  - Large db is needed with sufficient number of pathological cases and similar signal quality. Treatment of unbalance sets necessary to achieve results comparable to classification. (Janickova2014)



### Problems with goals (2)



#### Baby-centric: Related to delivery as a whole

- Always healthy baby => rate of CS close to 100%?
  - Not a reasonable thing, right? Not everybody can drive a tank...
- "Happy" baby & mother
  - Ideal, unfortunately very vague how to define such a state? 1990s with current prenatal treatment/diagnostics? Is it possible?
- Good pH (or any other "objective") value
  - The objective marker is not very specific when used for delivery outcome assessment. Too many influences from other clinical factors (e.g. sex of the new-born).

#### M.D. centric

Complementary goals to any above





## **Our conclusion from the intro qs**

- "Happy" baby is our goal
- Related clinical questions
  - To monitor or not to monitor (intensively)?
  - To use SoA supplementary methods e.g. STAN?
  - To simplify guidelines e.g. along 4-rules/states by Schifrin?
  - To find relevant clinical features (and combinations) to the outcome?

#### Features from CTG – just one input to more complex system

- Features focused on are decelerations and variability via scaling properties of the signal
- Composite outcome (Latent class modeling) is decided based on pH, Apgar, and subj. eval. of CTG
- ✤ Based on CTG class different features will be used for further analysis