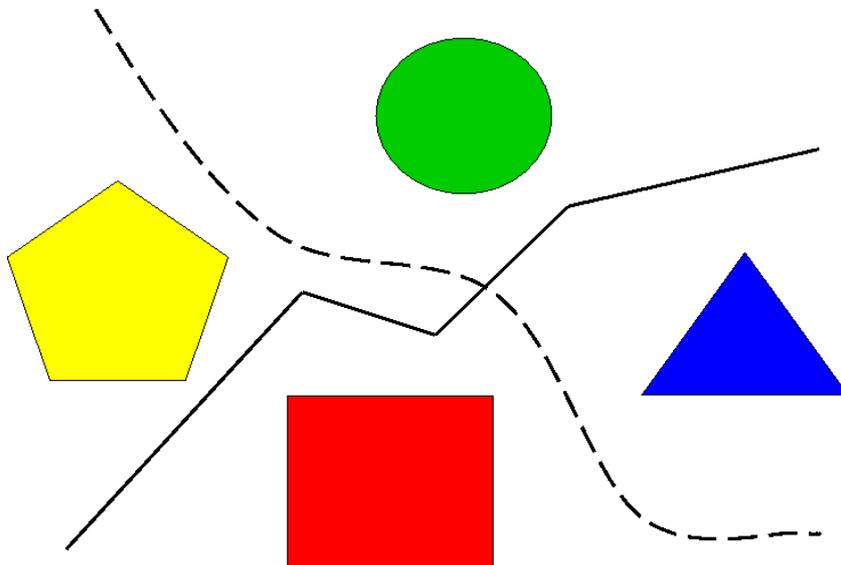


**6. Herbstkolloquium  
des Graduiertenkollegs  
"Statistische Modellbildung"**



**Statistische Modellbildung**

Zu diesem Kolloquium wird eingeladen

**Freitag / Samstag, 20./21. November 2009**

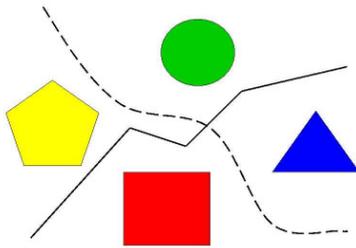
**UNIVERSITÄTSKOLLEG BOMMERHOLZ**

- Lehr- und Weiterbildungsstätte der Universität Dortmund -

**Bommerholzer Straße 60, 58456 Witten.**

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**Statistische Modellbildung**

## 6. Herbstkolloquium des Graduiertenkollegs "Statistische Modellbildung"

**Freitag, 20. November 2009**

**Abfahrt nach Witten: ab Dortmund gegen 14.00 Uhr**

### Vortragsprogramm I

- |              |  |   |
|--------------|--|---|
| <b>15:15</b> | <b>Begrüßung</b><br>Prof. Dr. Joachim KUNERT   |   |
| <b>15:30</b> | <b>Dr. Andreas FALDUM</b><br><i>IMBEI, Universitätsmedizin<br/>Johannes Gutenberg-Universität Mainz</i>  | <b>Strategies for including patients recruited during interim analysis of clinical trials</b> |
| <b>16:15</b> | <b>Dr. Harald BINDER</b><br><i>Institut für Medizinische Biometrie und<br/>Medizinische Informatik<br/>Universitätsklinikum Freiburg<br/>Freiburger Zentrum für Datenanalyse und<br/>Modellbildung<br/>Albert-Ludwigs-Universität Freiburg</i> | <b>Estimating regression models by likelihood-based boosting</b>                              |
| <b>17:00</b> | <b>Dr. Marco GRZEGORCZYK</b><br><i>Fakultät Statistik<br/>Technische Universität Dortmund</i>  | <b>Modelling non-stationary gene regulatory processes</b>                                     |

### Diskussion zu den Projektbereichen

- |              |  |
|--------------|--|
| <b>19:00</b> | <b>Posterausstellung:</b><br><b>Präsentation der Dissertationsprojekte im Kolleg, Diskussion in Arbeitsgruppen</b> |
|--------------|--|

**Samstag, 21. November 2009**

**Vortragsprogramm II**

**9:00**      **Dr. David GINSBOURGER**      **Towards Gaussian Process-based  
Optimization with Finite Time Horizon**  
*Centre d'Hydrogéologie et de Géothermie  
Université de Neuchâtel, CH*

**9:45**      **Dr. Olivier ROUSTANT**      **Computation of Sobol indices with a  
Gaussian Process metamodel**  
*Ecole des Mines, Département 3MI  
St.Etienne, F*

**10:30**      **Pause**

**Vortragsprogramm III**

**11:00**      **Prof. Dr. Dibyen MAJUMDAR**      **Optimal designs for two level factorial  
experiments with binary response**  
*Department of Mathematics, Statistics and  
Computer Science  
University of Illinois at Chicago, U.S.A.*

**11:45**      **Dr. Michael MEYERS**      **Statistical inference for Temporal Dominance  
of Sensations (TDS) data**  
*Nestlé Research Center  
Lausanne, CH*

**12:30**      **Pause**

**13:15**      **Gruppenfoto auf der Terrasse (wetterabhängig)**

**Vortragsprogramm IV**

**13:30**      **Prof. Dr. Anurag BANERJEE**      **Sensitivity of statistics against nuisance  
parameters: Measurement and solution**  
*Department of Economics and Finance  
Durham University, U.K.*

**14:15**      **Prof. Dr. Christian KLEIBER**      **A majorization approach to  
Condorcet jury theorems**  
*Wirtschaftswissenschaftliches Zentrum  
Universität Basel, CH*

**15:00**      **Prof. Dr. Rafael WEISSBACH**      **A Likelihood Ratio test for Stationarity of  
Rating transitions**  
*Wirtschafts- und Sozialwissenschaftliche  
Fakultät, Universität Rostock*

**15:30**      **Kaffeetrinken, Abschlussbesprechung und Diskussion**

**Abstracts**

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**Sensitivity of statistics against nuisance parameters: Measurement and Solution**

**Anurag Banerjee**

*Department of Economics and Finance  
Durham University, U.K.*

Econometric estimates or decisions are applicable if they are not sensitive to small changes of "nuisance" parameters. The statistic which measures these violations is called a sensitivity measure. We define such a measure and analyse its properties. If the measure suggests that when the sensitivity statistic is "small", we need not change the assumptions on the nuisance parameters. We also to propose a correction factor for the statistic of interest if we find "large" sensitivity. We shall define such an estimator and analyse its properties.

**Estimating regression models by likelihood-based boosting**

**Harald Binder (1,2)**

*1) Institut für Medizinische Biometrie und Medizinische Informatik, Universitätsklinikum Freiburg  
2) Freiburger Zentrum für Datenanalyse und Modellbildung, Albert-Ludwigs-Universität Freiburg*

Boosting techniques are based on the powerful idea of fitting several predictive models in a stepwise procedure, where each fit improves for observations that were fitted badly by the previous fits. An overall fit then is obtained by combining all fitted models. Given that each fit is additive in the covariate effects, the overall fit will be an additive regression model, providing simple structure. Likelihood-based boosting incorporates previous boosting steps as an offset into maximum likelihood estimation. Therefore, it is applicable to many classical regression models, allowing for estimation where standard maximum likelihood techniques fail. This will be illustrated using application examples with high-dimensional covariate vectors. Specifically, predictive survival models will be built from gene expression data. There, the proposed boosting techniques also allow to incorporate various sources of additional knowledge.

**Strategies for including patients recruited during interim analysis of clinical trials**

**Andreas Faldum**

*Universitätsmedizin der Johannes Gutenberg-Universität Mainz  
Institut für Medizinische Biometrie Epidemiologie und Informatik*

In clinical trials a periodical check of safety and efficacy data is often needed. For organizational reasons it is rarely desirable to stop a trial during such an interim analysis. Therefore, new study patients are included in the trial while the interim analysis is ongoing. Disregarding the additional information provided by these interim patients would be unsatisfactory, especially for an office of regulatory affairs. Consequently, the rules for group sequential or adaptive decisions must be adjusted to the recruitment of interim patients.

In the context of two prospective randomised adaptive trials, different modifications of the conditional error function are discussed to consider the analysis of interim patients. The impact of the proposed modifications on power and sample size is demonstrated. With an adequate consideration of interim patients the maximum sample size can be reduced considerably and a sufficient conditional power of the second stage is guaranteed.

## **Towards Gaussian Process-based Optimization with Finite Time Horizon**

**David Ginsbourger**

*Centre d'Hydrogéologie et de Géothermie  
Université de Neuchâtel, CH*

During the last decade, Kriging-based sequential algorithms like EGO and its variants have become reference optimization methods in computer experiments. Such algorithms rely on the iterative maximization of a sampling criterion, the expected improvement ( $EI$ ), which takes advantage of Kriging conditional distributions to make an explicit trade-off between promising and uncertain search space points. We have recently worked on a multipoints  $EI$  criterion meant to simultaneously choose several points, which is useful for instance in synchronous parallel computation. The research results that we wish to present on the occasion of this talk concern sequential procedures with a fixed number of iterations. We claim that maximizing the 1-point criterion at each iteration ( $EI$  algorithm) is suboptimal. In essence, the latter amounts indeed to considering the current iteration as the last one. During this talk, the problem of optimal strategy for finite horizon sequential optimization will be formulated, its solution will be provided in terms of multipoints  $EI$ , and the suboptimality of the usual  $EI$  algorithm will be illustrated on the basis of a first counter-example.

## **Modelling non-stationary gene regulatory processes**

**Marco Grzegorzcyk**

*Fakultät Statistik  
TU Dortmund*

One goal in systems biology is to infer regulatory networks from postgenomic data. Bayesian networks have been widely applied as a popular tool to this end. To obtain a closed-form expression of the marginal likelihood two probabilistic models with their respective conjugate prior distributions have been employed: the multinomial BDe and the Gaussian BGe scoring metric. These scores are restricted in that they either require a data discretisation (BDe) or can only capture linear relationships (BGe). In an earlier work we proposed a generalisation of BGe based on a combination of a mixture model and the allocation sampler. In the BGM model data points are allocated to different compartments (subsets) by a free individual allocation sampler, and the same single network structure is inferred for all compartments. Given the network structure each compartment is modelled separately and independently with the Gaussian BGe model. The fixation of the network allows for some information sharing among compartments. This work specialises the BGM model with respect to dynamic gene regulatory networks by replacing the free allocation model by a changepoint process that takes the time structure into account. We also show that discrete counterparts of the free allocation model BGM and the novel changepoint process model BGMD, which employ the multinomial BDe score for modelling the data compartments, can be used for inferring non-stationary regulatory processes from discrete expression data.

**Abstracts**

**A majorization approach to Condorcet jury theorems**

**Christian Kleiber**

*Wirtschaftswissenschaftliches Zentrum  
Universität Basel, CH*

The partial order of majorization is omnipresent in applied mathematics, statistics and various fields of application. It suggests comparing two given vectors, for example representing the incomes of two populations, by comparing the partial sums of their ordered entries. Among other applications, majorization can be used to study probability inequalities for sums of heterogeneous Bernoulli variables. These arise in the context of the 'Condorcet jury theorem', a political science theorem about the relative probability of a given group of individuals arriving at a correct decision. This talk shows how majorization inequalities can shed new light on Condorcet jury theorems for heterogeneous juries.

**Optimal designs for two level factorial experiments with binary response**

**Dibyen Majumdar**

*Department of Mathematics, Statistics and Computer Science  
University of Illinois at Chicago, U.S.A.*

We consider the problem of finding locally D-optimal designs for two level factorial experiments with binary response using a generalized linear model. The optimal designs are obtained using analytic and computational methods. The optimal designs depend on the initial values of the parameters and the choice of the link function. The robustness of the designs is studied. In this talk we will focus on the case of two factors. This is ongoing work with Abhyuday Mandal and Jie Yang.

**Statistical inference for Temporal Dominance of Sensations (TDS) data**

**Michael Meyners**

*Nestlé Research Center  
Lausanne, CH*

Temporal dominance of sensations (TDS) is a recently developed sensory test procedure. Throughout a certain period, subjects are asked to judge continuously which out of a few attributes (up to 8, say) is the dominant one, and also to rate the intensity of the dominant attribute. Usually considered periods of interest include the mastication period and the aftertaste period.

Up to now, statistical analysis is confined to a description of the data, the depiction of the average dominance curves, and some rough cut-off limits based on the binomial law. We propose an appropriate and valid set of statistical tests that allows the investigation of overall differences as well as of differences between pairs of products. Next to a general test, inferential methods per attribute, per time point and per attribute by time point are suggested. All tests are based on the general notion of randomization tests (cf. Edgington & Onghena, 2007), and they are valid level- $\alpha$  tests by themselves. Without correction for multiplicity, the set of tests of course does not respect the family-wise significance level. Strict corrections (like Hommel's, 1988) or corrections based on the false discovery rate (Benjamini & Hochberg, 1995) can be applied to the findings to reduce the risk or incidence rate of false positive results.

To start the presentation, the concept of TDS will be introduced. We then motivate the assumption that the time periods are standardized across subjects and products, such that each observation has a fixed number of time

**Abstracts**

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points. The impact of this assumption will be briefly discussed. Furthermore, we confine ourselves to the pure choice of attributes and neglect the intensity scores for the remainder, but will discuss some generalizations in this regard towards the end of the presentation.

We describe how the TDS sequences are unfolded to data matrices with a single non-zero entry per time point (column). The sum of the Euclidean distances between these matrices is determined and serves as a test statistic for the global test. Similar statistics can be used for pairwise comparisons and for inference by attribute or time point. Re-randomizations are used to determine the null distribution and p values for each hypothesis under consideration, taking the original restrictions of the randomization into account. We also propose a simple graphical method to summarize the many p values derived from this approach (usually at least hundreds, but more often several thousands). An application to real data is shown to give reasonable and easily interpretable results.

- Benjamini, Y., & Hochberg, Y. (1995). Controlling the false discovery rate: a practical and powerful approach to multiple testing. *Journal of the Royal Statistical Society Series B*, **57**, 289–300.
- Edgington E, & Onghena P (2007). *Randomization tests*. 4<sup>th</sup> ed., Chapman & Hall/CRC.
- Hommel, G. (1988). A stagewise rejective multiple test procedure based on a modified Bonferroni test. *Biometrika*, **75**, 383–386.

## **Computation of Sobol indices with a Gaussian Process metamodel**

**Olivier Roustant**

*Ecole des Mines, Département 3MI  
St-Etienne, F*

When using time expensive computer codes, performing a global sensitivity analysis directly on the code may be untractable. To overcome that problem, an approach is to replace the code by a quick-to-evaluate metamodel. In this talk we focus on the famous Gaussian process (GP) model, coming from kriging, and address some computations issues. First, when replacing the code by the GP mean, we explain how to obtain analytical expressions of Sobol indices, under suitable assumptions. We give the computation details for the indices corresponding to main effects and interactions. In the second part, we discuss some extensions - what's happening when adding a linear trend to the model? When replacing the code by the global GP (and not only its mean)? - and some practical limitations.

## **A Likelihood Ratio Test for Stationarity of Rating Transitions**

**Rafael Weißbach**

*Wirtschafts- und Sozialwissenschaftliche Fakultät  
Universität Rostock*

We study the time-stationarity of rating transitions, modelled by a time-continuous discrete-state Markov process and derive a likelihood ratio test. For multiple Markov processes from a multiplicative intensity model, maximum likelihood parameter estimates can be written as martingale transform of the processes, counting transitions between the rating states, so that the profile partial likelihood ratio is asymptotically chi-square-distributed. An application to an internal rating data set reveals highly significant instationarity.