Saarland University Center for Bioinformatics Master's Program in Bioinformatics



Master Thesis in Bioinformatics

$\begin{array}{c} \textbf{Design and calibration of stochastic models for} \\ \textbf{DNA methylation patterns} \end{array}$

submitted by

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on March 2019

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Design and calibration of stochastic models for DNA methylation patterns

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Declaration

I hereby confirm that this thesis is my own work and that I have documented all sources used.

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Saarbrücken, on March 2019

Andrea Kupitz

Abstract

The expression of genes in the human genome is not only based on the DNA sequence; it relies on epigenetic modifications like DNA-methylations. Hereby, gene expression is inactivated by binding of methyl-groups to a cytosine phosphate guanine (CpG) dinucleotide at the promoter region of the concerned gene. The binding is performed by specific enzymes - the DNA Methyltransferases (DNMTs). The specific function of DNMTs is not fully determined.

In the following, the methylation of DNA is modelled using an Markov Chain Monte Carlo (MCMC) algorithm and parameters are estimated by Maximum Likelihood Estimation (MLE). Alternatively, parameter estimation is performed with an implementation of the Approximative Bayesian Computation (ABC) method. Moreover, a method to compare distributions of methylation patterns is provided.

We find differences in the function of the different DNMTs that are consistent with current biological findings. Thus, using this model, some parameters are difficult to identify because they seem to be conditionally dependant.

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ABC	Approximative Bayesian Computation	ii
\mathbf{CpG}	cytosine phosphate guanine	ii
$\overline{ ext{DNMTs}}$	DNA Methyltransferases	i
MCMC	Markov Chain Monte Carlo	i
MLE	Maximum Likelihood Estimation	ii



Introduction

Methods

Evaluation

Discussion

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Appendix A

Regulation