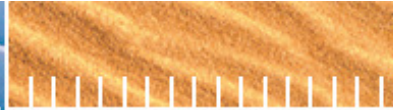




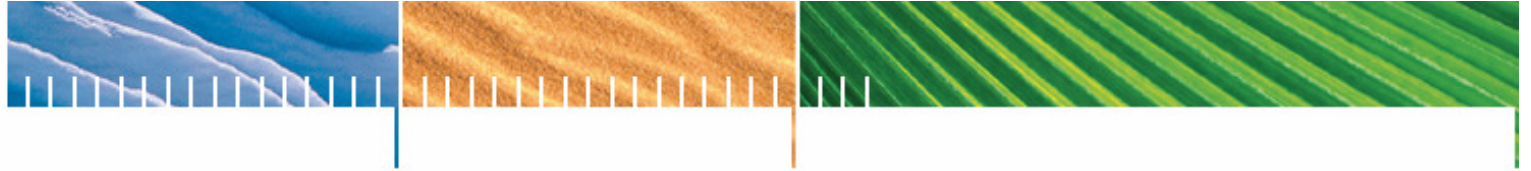
Swiss Tropical Institute
Institut Tropical Suisse
Schweizerisches Tropeninstitut



Estimating parameters for mathematical models of malaria epidemiology

Nicolas Maire
Swiss Tropical Institute
malariacontrol.net
BOINC Workshop 09



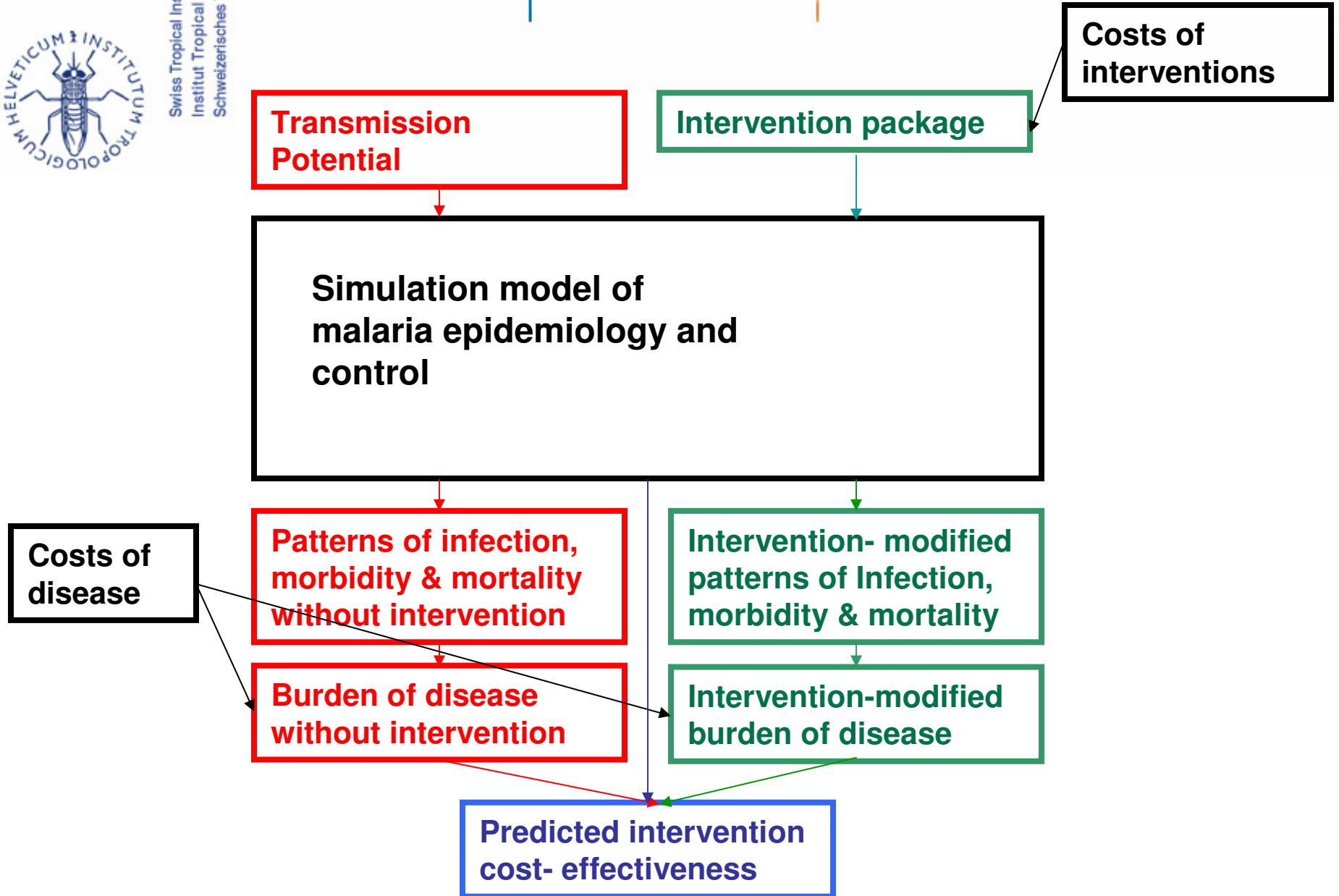
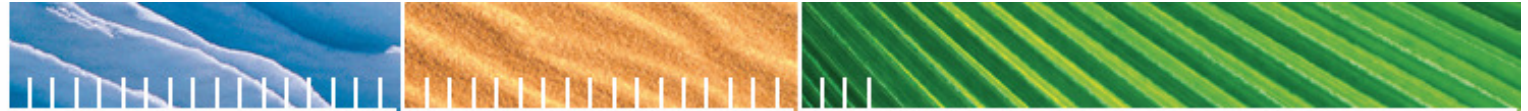


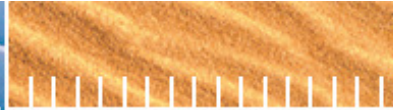
Malaria and malaria models

- Malaria: a mosquito-borne disease caused by a parasite
- One to three million deaths per year
- Hundreds of millions of illness episodes per year, up to 40% of health expenditure
- Mathematical models have been a valuable decision making tool in public health
- Increasing demand for rational setting of priorities



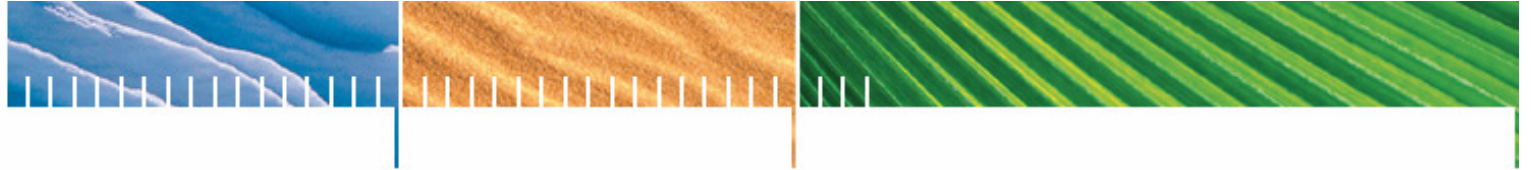
Swiss Tropical Institute
Institut Tropical Suisse
Schweizerisches Tropeninstitut



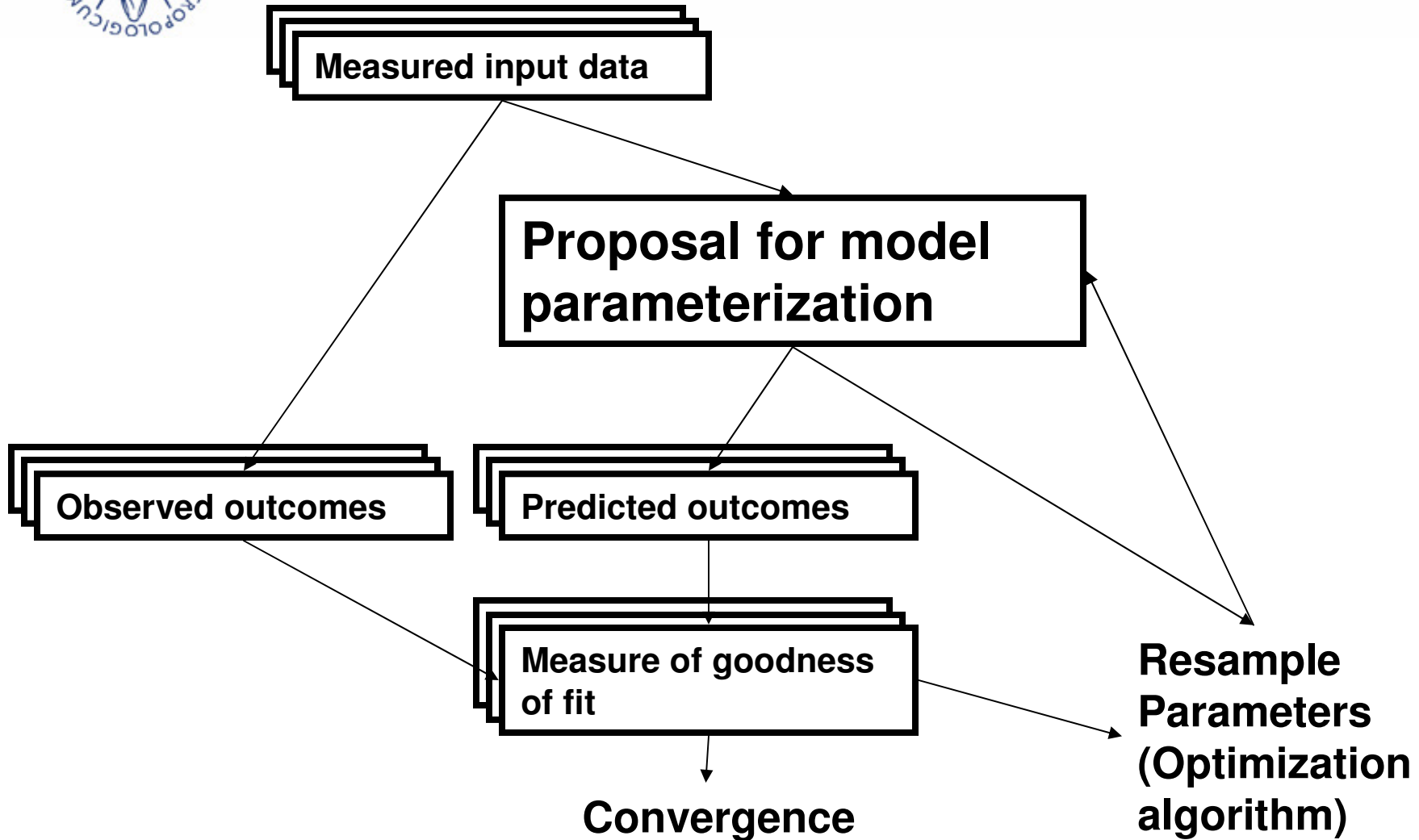


Modeling approach

- Discrete time stochastic individual-based simulations
 - Human hosts are characterized by a set of state variables (age, parasitology, immune status,...)
- Models for the effect of acquired immunity on parasite densities, for transmission to the vector, for morbidity, and for mortality
- Fit model to data from field studies
- Predict impact of control strategies by comparing simulated interventions with baseline scenarios



Estimating model parameters from field data





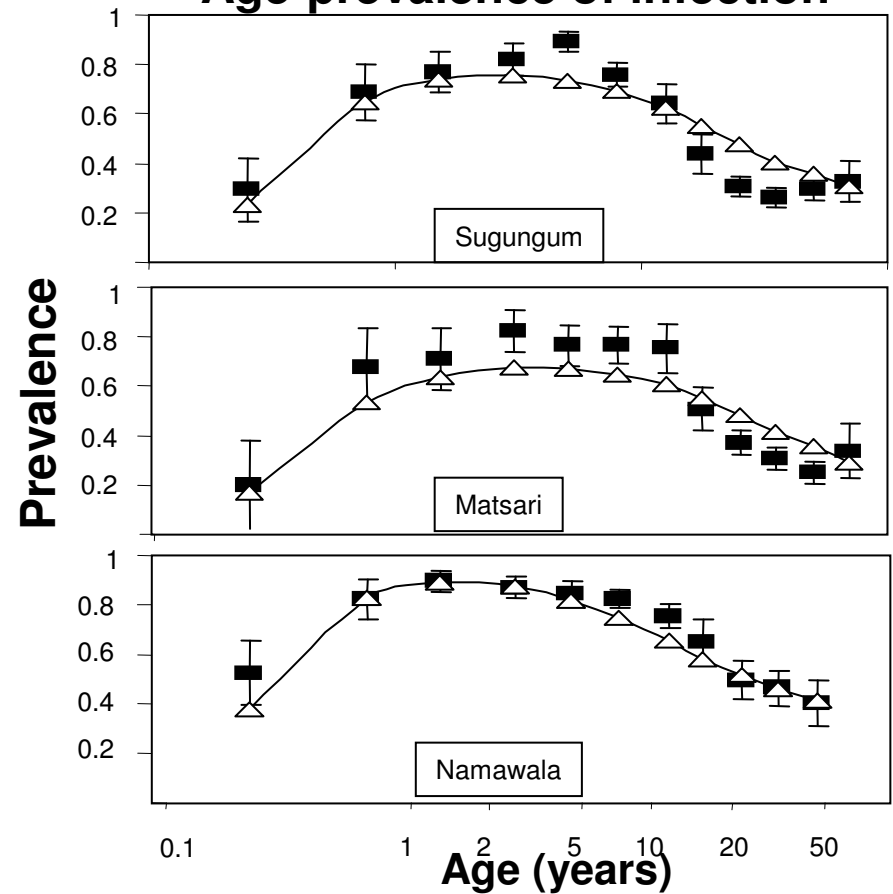
Swiss Tropical Institute
 Institut Tropical Suisse
 Schweizerisches Tropeninstitut



Calibration

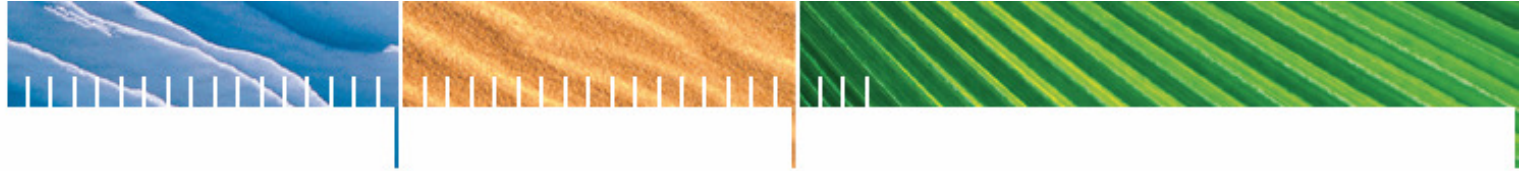
- 61 datasets from field studies, different objectives*
 - Incidence of infection
 - Age-prevalence of parasitemia
 - Seasonality of parasitemia
 - Age-density of parasites
 - Age-incidence of clinical disease, hospitalisation and mortality

Age-prevalence of infection



■ Field data (CI) △ Model

*all related to seasonal patterns of transmission



Challenge of fitting these models

- Objective functions non-differentiable
- Loss function values are not reproducible because of stochasticity
- High-dimensional parameter space
- (Multi-objective)
- Computationally expensive
- *malariacontrol.net*
- Time to completion of results long and unpredictable

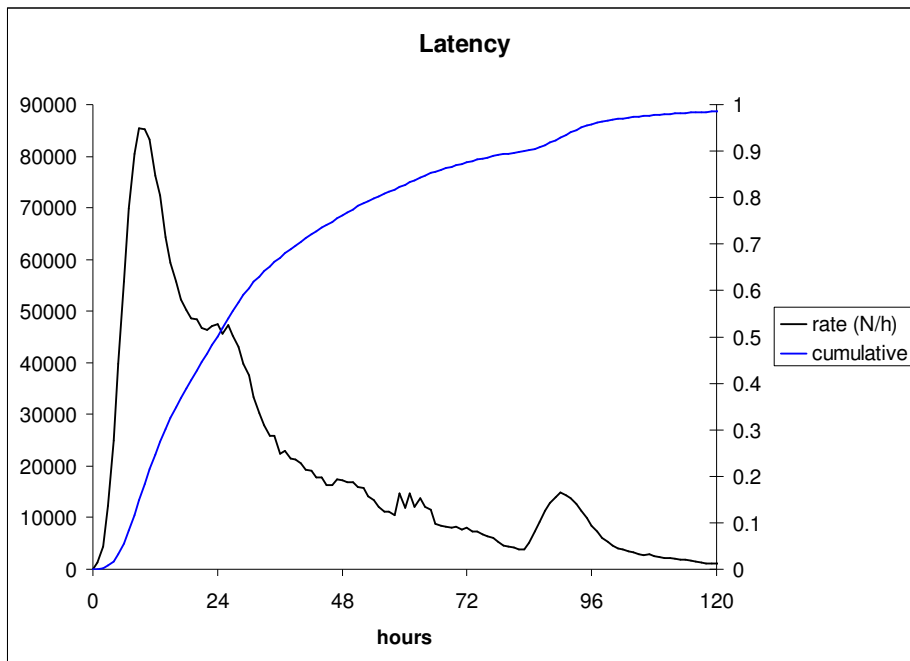


Swiss Tropical Institute
Institut Tropical Suisse
Schweizerisches Tropeninstitut

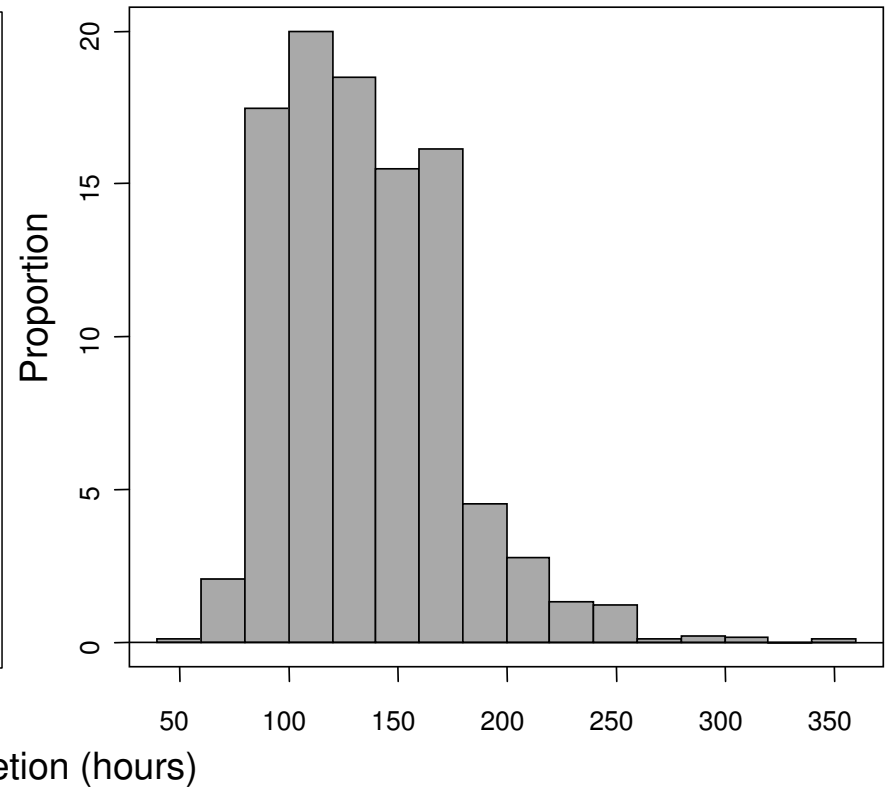


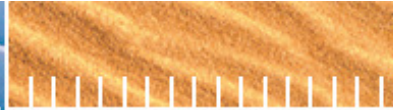
Latency

Workunits



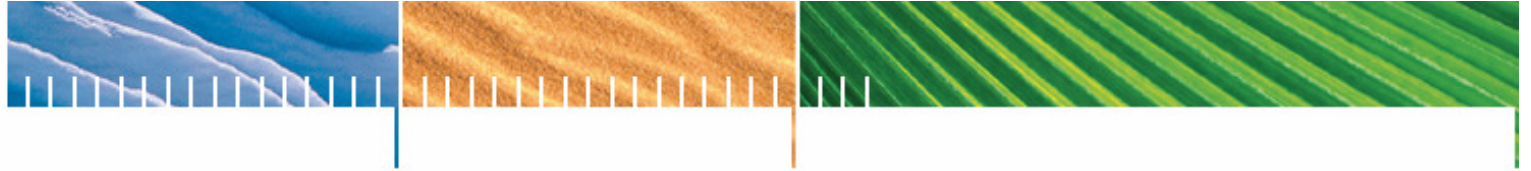
Parameterizations





BOINC Scheduler improvements

- Reliable host scheduling
- Redundancy elimination (next release of mcdn science app)

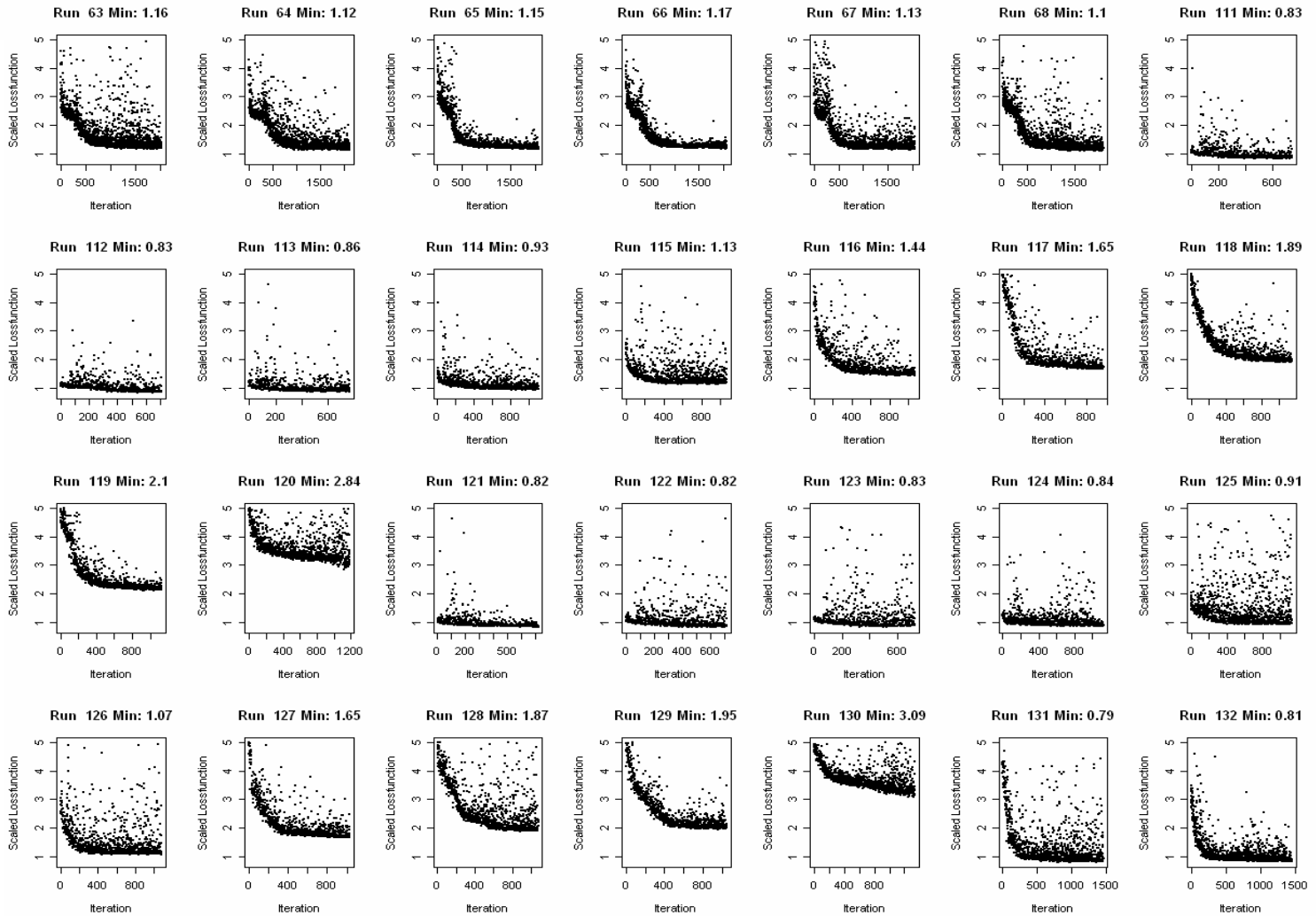


Current strategy using Genetic Algorithm

- Truncating mating selection
 - Choose best n individuals
- Mutation
 - Sample around current parameter values
 - Adaptive sampling range
 - Adaptive mutation rate
- Overlapping generations
 - Work generator samples new parameterizations on demand

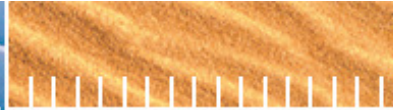


Fit by iteration for alternative models



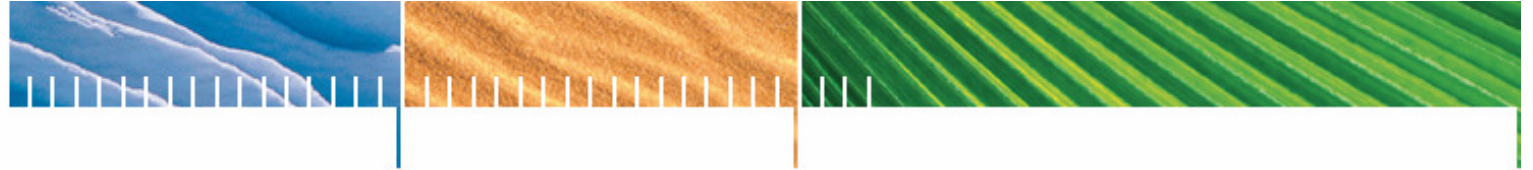


Swiss Tropical Institute
Institut Tropical Suisse
Schweizerisches Tropeninstitut



Questions

- How much do we gain by increasing the number of participating hosts?
- Which optimization algorithm works best?

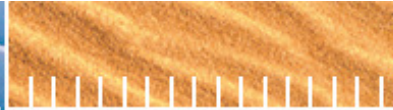


Return on investment in additional hosts

- Simulate different volunteer population sizes by using different workunit generation priorities
- Compare optimization performance against number of participating hosts
- Metrics
 - Performance: Absolute time to convergence
 - Investment: Avg. number of parameterizations sampled per evaluation period

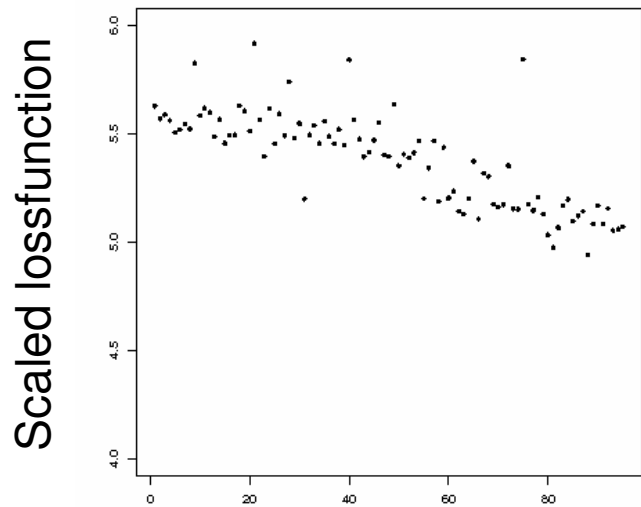


Swiss Tropical Institute
Institut Tropical Suisse
Schweizerisches Tropeninstitut

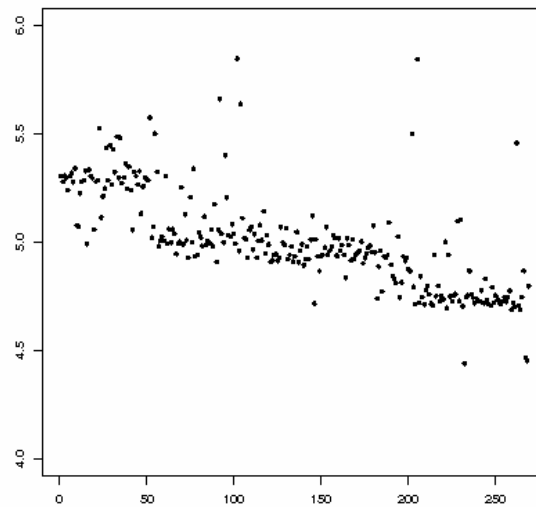


Preliminary results

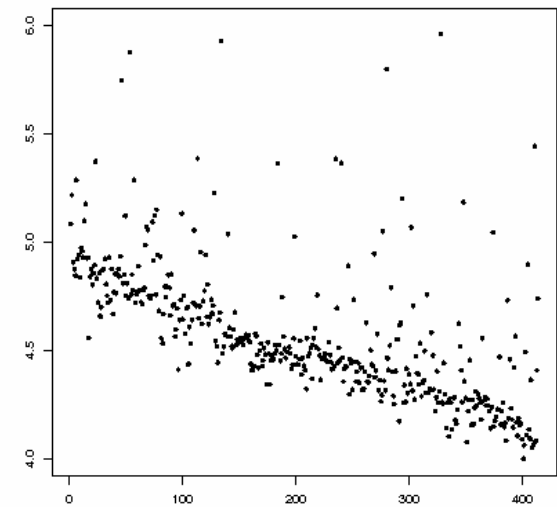
18 samples/eval



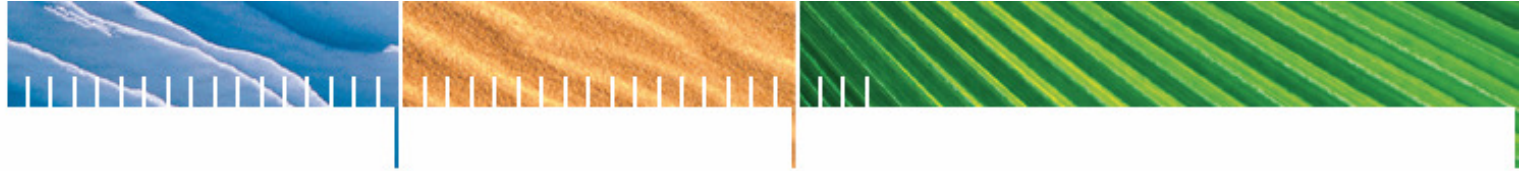
36 samples/eval



72 samples/eval



Iteration/Time

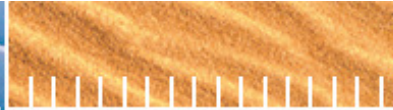


Comparison of different optimization algorithms

- Design work generator for pluggable algorithms
- Mostly a wrapper for the backend database
- Abstract **Algorithm** class with sample method
 - `Parameterization sample(Population pop);`
- Core classes
 - Population
 - Parameterization

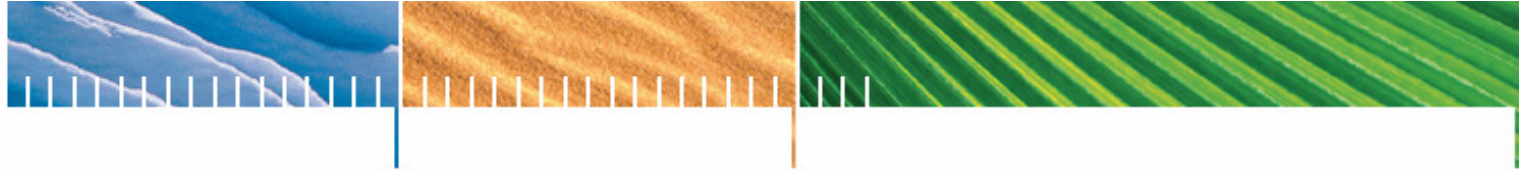


Swiss Tropical Institute
Institut Tropical Suisse
Schweizerisches Tropeninstitut



Population

- A collection of completed Parameterizations
 - `vector<Parameterization> getParameterizations();`
 - `string getProperties();`
 - `string setProperties();`

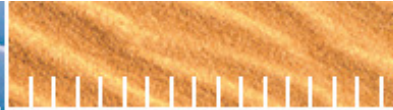


Parameterization

- A collection of samplable parameters, a lossfunction, and optional properties
- `int getNumParameters();`
- `double getLossfunction();`
- `double getValue(int index);`
- `void setValue(int index, double value);`
- `string getProperties();`
- `void setProperties(string properties);`



Swiss Tropical Institute
 Institut Tropical Suisse
 Schweizerisches Tropeninstitut



Current Research Team

Applied Mathematics

Melissa Penny (STI)
 Nakul Chitnis
 (STI/MACEPA)

Epidem./Public Health

Allan Schapira (STI)
 Blaise Genton (STI)
 Christian Lengeler (STI)
 Don de Savigny (STI)
Marcel Tanner (STI)

Quantitative biology

Ian Hastings (LSTM)
 Katherine Winter (LSTM)
 Michael Bretscher (STI)

Statistics

Amanda Ross (STI)
Tom Smith (STI)

Computer Science

Diggory Hardy (STI)
 Aurelio di Pasquale (STI)
 Nicolas Maire (STI)
 Tiago Antão (LSTM)
 Henning Mortveit (VBI)

Health Economics

Fabrizio Tediosi (Milan)
 Josh Yukich (STI)
 Lesong Conteh (LSHTM)
 Valerie Crowell (STI)

Spatial Statistics

Penelope Vounatsou (STI)
 Laura Gosoni (STI)
 Nadine Riedel (STI)
 Amina Msengwa (STI)

Databases

Konstantina Boutsika (STI)
 Tanja Jäggi (STI)
 Eric Diboulo (Ouaga/STI)

Financial support

Bill & Melinda Gates Foundation, PATH-MACEPA, Swiss National Science Foundation