

## ArtiSaneFood Annual Meeting

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Bologna, Italy, 7-8 July, 2022

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## Context of my thesis

- **ArtisaneFood France**: Study impact of intervention strategies (WP 7)
  - Product: Camembert de Normandie (fromage au lait cru)
  - Pathogen: Shiga Toxin producing Escherichia coli (STEC)
  - Disease: Haemolytic Uremic Syndrome (HUS)
- **Thesis duration**: January 2021 - December 2023
- **Academic advisors**:
  - Julien BECT (L2S) & Emmanuel VAZQUEZ (L2S)
  - Laurent GUILLIER (ANSES)
- **Industrial advisors**:
  - Fanny TENENHAUS AZIZA (CNIEL)



This work is part of the **ArtiSaneFood** project (grant number : **ANR-18-PRIM-0015**) which is part of the **PRIMA** program supported by the **European Union**.

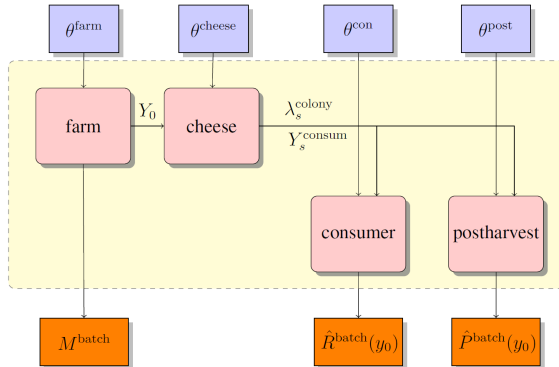


# Contents

- 1 Building QRA simulator**
- 2 Publishing the QRA model**
- 3 Study impact of intervention strategies**
- 4 Work in progress!**

# 1 Building QRA simulator

- QRA model proposed by [Perrin et al. \(2014\)](#)



## Batch level simulator

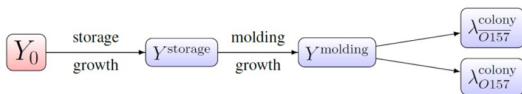
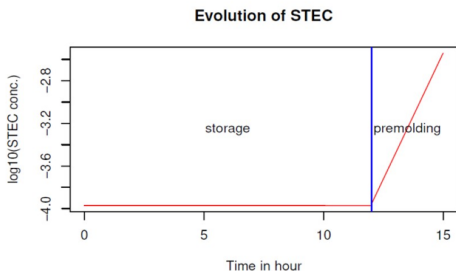
- Implemented in **R**
- Models **farm-to-fork** continuum
- Simulates fabrication of one cheese batch
  - Contains 22,000 to 23,000 cheese (250 g)
- **Outputs:**
  - Milk loss due to farm rejection
  - Risk of HUS from MPS-STEC
  - Probability of batch rejection

## The modules

- Farm module
  - Collection of milk from farms
  - Pre-harvest intervention (milk testing): Test E.Coli conc.
  - **Outputs**: STEC concentration (CFU/ml) in milk to be used
- Cheese module
  - Evolution of STEC using ODEs
  - STEC cells form clusters (a.k.a colonies) in cheese
  - **Outputs**
    - \* Number of colonies in CFU (Poisson)
    - \* Size of colonies (LogNormal)

## Evolution of STEC (liquid phase)

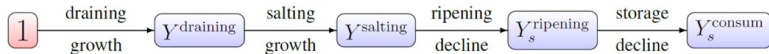
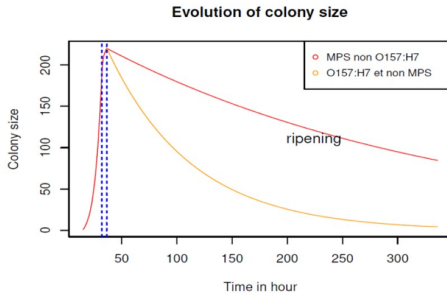
$$\frac{dy}{dt} = \mu^{\max}(t) \cdot y(t) \cdot \left(1 - \frac{y(t)}{y^{\max}}\right)$$





## Evolution of STEC (solid phase)

$$Y_s^{\text{consum}} = Y_s^{\text{salting}} \cdot 10^{-\rho_s \cdot (t^{\text{consum}} \times 24 - t) / 24}$$



## The modules

- Consumer module

- Risk is computed using a dose response model

$$\Gamma = \sum_s N_{s,\text{sample}}^{\text{colony}} \cdot Y_s^{\text{colony}}$$

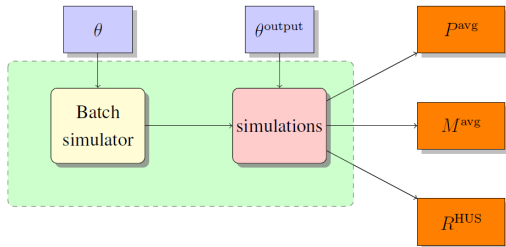
- Averaged over consumption behaviour for different ages
- **Outputs** Risk of HUS from MPS-STECS

- Post-harvest module

- Cheese batches are tested for STEC contamination
- **Outputs**
  - \* Probability of rejecting a batch

## Final quantities of interest

- Several batches are simulated



- Overall risk of HUS
- Average milk loss
- Proportion of rejected batches

## Modification & improvements

- **Bayesian approach** to estimate E.Coli distribution parameters
  - Using E.Coli concentration data from CNIEL
- **Adaptive algorithm** to reduce the computational time
  - **Publication** : S. Basak, J. Bect , and E. Vazquez. Integration of bounded monotone functions: Revisiting the nonsequential case, with a focus on unbiased Monte Carlo (randomized) methods. In *53èmes Journées de Statistique de la SFdS, Lyon, France, June 2022*
- **Analytical solution** for estimating batch rejection probabilities
  - Replacing Negative Binomial distributional assumption on dose
- ...

## 2 Publishing the QRA model

- Food and Ecological Systems Modelling Journal [FESMJ](#)
  - Article with description/functionalities of the model (**WIP!**)
- Food Safety Knowledge Exchange [FSKX](#)
  - A FSKX version of the model is available
- [RAKIP](#) repository
  - The model will be available on this repository



### 3 Study impact of intervention strategies

- The QRA simulator is **stochastic** + **expensive**
- **Pre-harvest** intervention (milk testing) parameters
  - Frequency of milk testing
  - Test threshold of E.Coli in CFU/ml
- **Post-harvest** intervention (cheese testing) parameters
  - Proportion of batches tested
  - Number of cheese samples tested
- **Objectives to minimize**
  - Risk of HUS
  - Cost due to rejecting milk & cheese batches

## Multiobjective optimization

- We consider a biobjective optimization problem

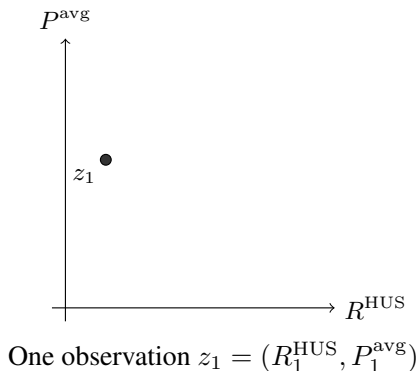
$$\min_{x \in \mathbb{X} \subset \mathbb{R}^2} f(x) \quad (1)$$

- The solution set does **not** contain a unique optimal solution
- It contains **Pareto optimal** points

$$\mathcal{P} = \{x \in \mathbb{X} : \nexists x' \in \mathbb{X}, f(x') \prec f(x)\} \quad (2)$$

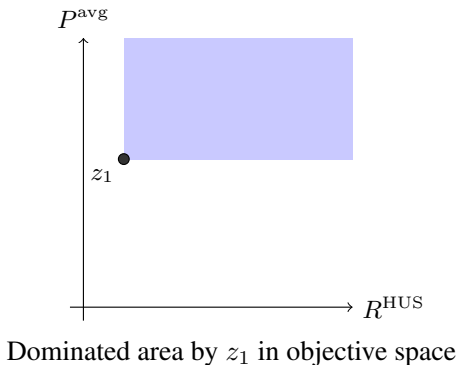
- Where  $f' \prec f \implies f'_i \leq f_i, \forall i$ , with at least one of the inequalities being strict

## Pareto optimal solutions: the objective space

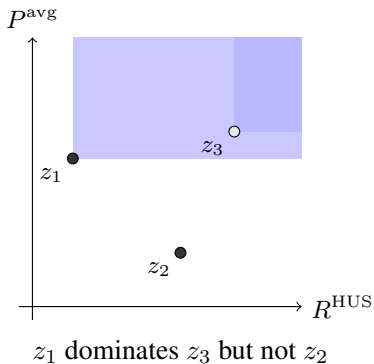




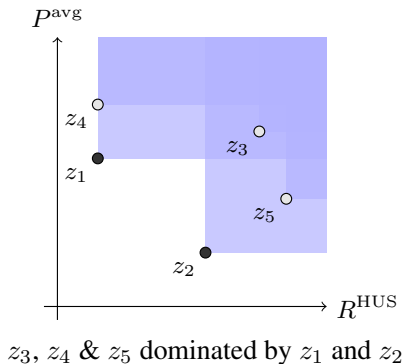
## Pareto optimal solutions



## Pareto optimal solutions



## An illustration: Pareto optimal solutions

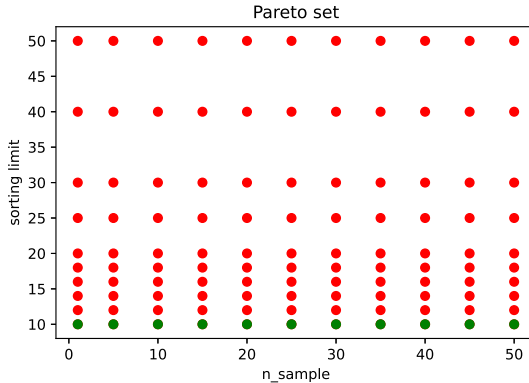


## The optimization problem: A first approach

- Minimize the function  $f = (R^{\text{HUS}}, P^{\text{avg}})$
- To find: **optimal** value of  $(l^{\text{sort}}, n_{\text{sample}}) \in \mathbb{X}$  that minimize  $f$ 
  - **Experimental design**  $(l^{\text{sort}}, n_{\text{sample}}) \in \mathbb{X}$   
 $\mathbb{X} = \{1, 5, 10, 15, \dots, 50\} \times \{10, 12, 14, 16, 18, 20, 25, 30, 40, 50\}$
  - **Objective space**  $[0, 1]^2$
- **A naive solution**: Evaluate the simulator  $\forall x \in \mathbb{X}$ 
  - The simulator is **expensive**
  - The simulator is **stochastic**
- Therefore, we rely on Bayesian optimization algorithms

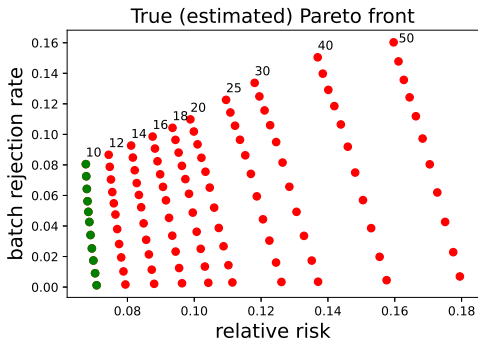
# True Pareto set (Experimental design)

- Pareto optimal (green) and non Pareto optimal (red)



## True Pareto front (Objective space)

- A heavy Monte-Carlo on  $\mathbb{X}$  (110 points take 9 hours!)
- Numbers denoting **sorting limit**  $l^{\text{sort}}$

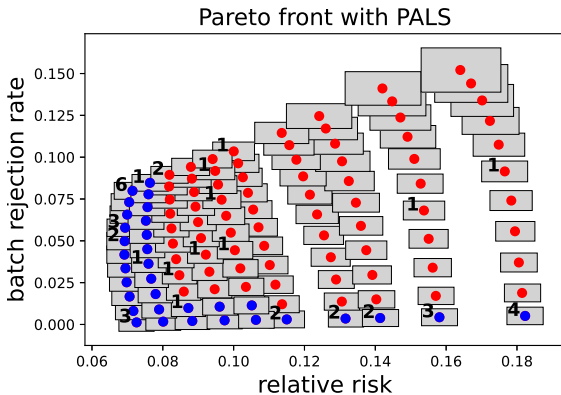


# Stochastic Pareto Active Learning (PALS)

- Easy to implement and inexpensive
- Proposed by [Zuluaga et al. \(2013\)](#) and extended by [Barracosa et al. \(2021\)](#)
- [Gaussian process](#) (GP) regression is used for surrogate modelling
  - Input space is classified by [confidence rectangles](#) using GP posterior mean and variance
- **Poster** : Subhasish Basak, Julien Bect , Laurent Guillier , Fanny Tenenhaus Aziza, Janushan Christy, Emmanuel Vazquez. Bayesian multiobjective optimization for quantitative risk assessment in microbiology. In *PhD students day in the Annual meeting of GdR MASCOT-NUM research Network, June 2022, Clermont Ferrand, France.*

## Estimated Pareto front with PALS

- Pareto optimal (green), unclassified (blue) and non Pareto optimal (red)





## 4 Work in progress!

- Extending PALS for
  - Grid-free continuous input space
  - more generalized confidence region
- Consider the correlated noise between Qols
- Consider other objectives like cost of **milk loss**, **batch loss**
- Integrate other design variables
  - Proportion of **milk** batch tested
  - Proportion of **cheese** batch tested

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## References

- F. Perrin, F. Tenenhaus-Aziza, V. Michel, S. Miszczycha, N. Bel, and M. Sanaa. Quantitative risk assessment of haemolytic and uremic syndrome linked to O157:H7 and non-O157:H7 shiga-toxin producing escherichia coli strains in raw milk soft cheeses. Risk Analysis, 35(1): 109–128, 2014.
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- B. Barracosa, J. Bect, H. Dutrieux Baraffe, J. Morin, J. Fournel, and E. Vazquez. Extension of the Pareto Active Learning method to multi-objective optimization for stochastic simulators. In SIAM Conference on Computational Science and Engineering (CSE21), Virtual Conference originally scheduled in Fort Worth, Texas, United States, Mar 2021.