

# A preliminary study of the para-model control of HIV

- Controlling without knowing -

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Friday 11<sup>th</sup> May, 2018



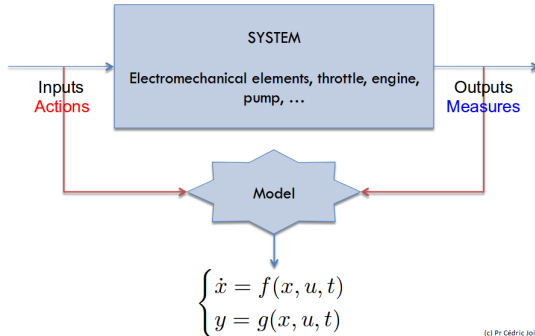
# OUTLINE

- Presentation of the model-free control methodology
- Toward Para-Model control
- A preliminary study of the para-model of the HIV-1

# Main idea of model-free control

Control today

- Most existing works: need precise mathematical (often difficult) modelling



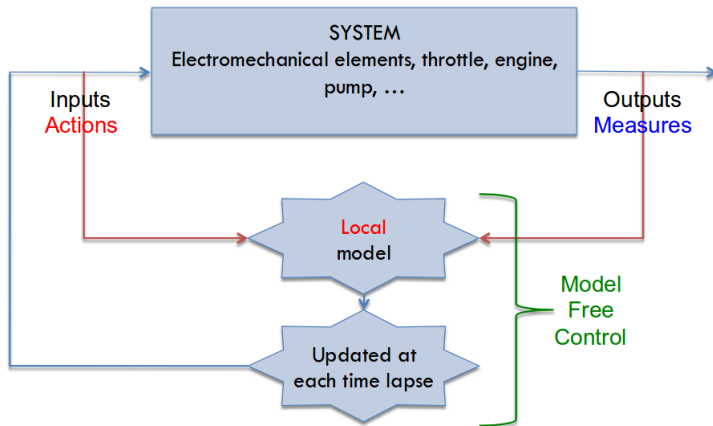
(c) Pr Cédric Join

- PID (Proportional Integral Derivative) most common control law in practice (>95 %)

# Main idea of model-free control

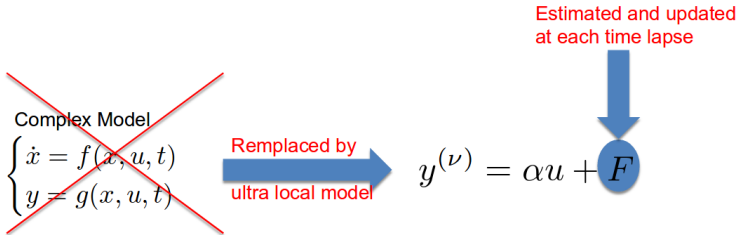
Model-free based control architecture

- From a complex system... to a simple local approximation



# Main idea of model-free control

## Model-free based control overview



Ultra local model control based

$$u = \frac{1}{\alpha} \left( -F + y_*^{(\nu)} \right) + \textit{Correction}(e)$$

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- Same conceptual simplicity as PID controllers
- No need of precise modelling

# Model-free control

Model-free based control discrete formal definition

**(Fliess, Join - 2008)**

Consider an unknown system  $E : u \mapsto y$ ,

$$E(t, v, \dot{v}, \dots, v^{(\iota)}, u, \dot{u}, \dots, u^{(\kappa)}) = 0$$

## MFC definition

To control  $y$  relating to a reference  $y^*$ , one considers the intelligent PI (i-PI) controller that reads:

$$u_k = u_{k-1} - \frac{1}{\beta} \left( \left. \frac{dy}{dt} \right|_{k-1} - \left. \frac{dy^*}{dt} \right|_k \right) + \mathcal{C}(y^*|_k - y|_{k-1})$$

where  $\mathcal{C}$  is a PI controller and  $\beta$  is a real constant; the tracking error is  $\varepsilon = y^*|_k - y|_{k-1}$

- Numerical derivation of  $y$  necessary

# Model-free control

In general, performances increase:

- development costs decrease
- maintenance costs decrease
- energetic costs decrease

Towards recent first applications in biology...

- Acute inflammation
- Wastewater denitrification
- Dynamic compensation for homeostasis
- Glycemia regulation

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# Para-model control

Based on the original Model-free control...

- Simplified version of the original model-free control

## Para-model control definition

To control  $y$  relating to a reference  $y^*$ , one considers the  $\mathcal{C}_\pi$  controller that reads:

$$u_k = \int_0^t K_i \varepsilon_{k-1} d\tau \Big|_{k-1} \left\{ u_{k-1} + K_p (k_\alpha e^{-k_\beta k} - y_{k-1}) \right\}$$

where  $K_i, K_p, k_\alpha, k_\beta$  are real coefficients to adjust

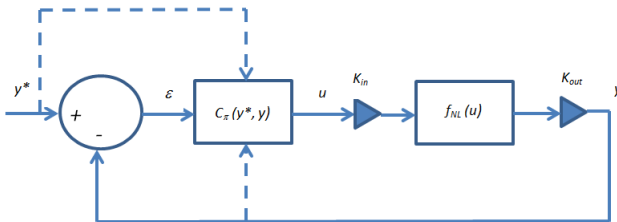
- Derivative-free algorithm

# Para-model control

## Structural properties

Given an output reference  $y^*$  and a nonlinear system  $y = f_{nl}(u)$ , it is *a priori* possible:

- to *control* dynamical or static  $f_{nl}$  systems
- to *optimize* (look for extremum) dynamical or static  $f_{nl}$  systems



# Para-model control

Matlab code

```
y_int(i) = M_alpha*exp(-M_beta*tt(i));  
para_exp_err = y_int(i-1) - y(i-1);  
para_stand_err(i) = y_ref(i) - y(i-1);  
para_u(i) = para_u(i-1) + Kp*para_exp_err;  
para_G(i) = Kint*para_stand_err(i);  
para_tr(i) = para_tr(i-1) + h*(para_G(i) + para_G(i-1))/2;  
para_u_final = para_u(i)*para_tr(i);
```

# BFO-tuning of the Para-model control

Tuning of the  $\mathcal{C}_\pi$  coefficients in simulation

(Porcelli, Toint - 2015)

- DFO-based Brute Force Optimization (BFO) Matlab package
- For a given closed-loop, consider a performance index  $\mathbb{P}$  (IAE, ISE, ITAE or ITSE) to minimize

## PMA Tuning Optimization Procedure

Consider  $\mathcal{C}_\pi$  that controls an unknown system  $E$  over  $[0, t]$ . We expect to minimize a performance index  $\mathbb{P}$  of the closed-loop:

$$\min_{K_i, K_p, k_\alpha, k_\beta} \mathbb{P} \quad \text{e.g.} \quad \min_{K_i, K_p, k_\alpha, k_\beta} \int_0^t (y - y^*)^2 dt \quad (\text{ISE})$$

# BFO-tuning of the Para-model control

## An example

- Consider a first order  $E$  dynamical system
- We expect to minimize  $\mathbb{P} = IAE + ISE + ITAE + ITSE$  over  $[0, t]$

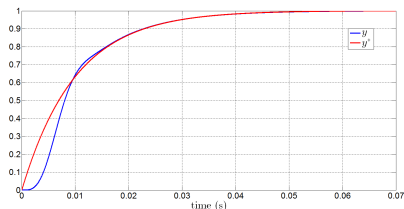


Figure:  $\mathcal{C}_\pi$  control of  $E$  without optimization

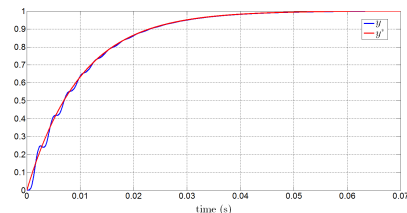


Figure:  $\mathcal{C}_\pi$  control of  $E$  with optimization

# First applications in simulation

## Présentation

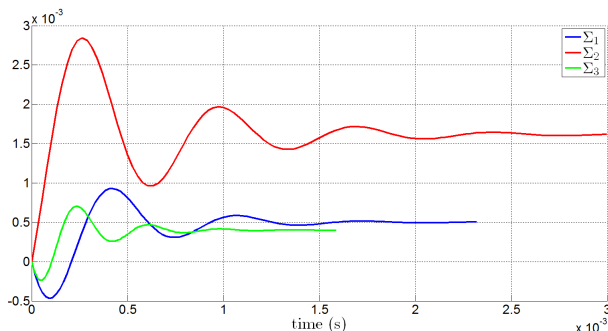
### *Application example:*

- Power electronics : control of nonlinear converters
- Mechanics : ballistic fire and robot trajectory
- Magnetism : characterization of magnetic materials (experimentally validated)
- Optimization : "extremum seeking-control"
- Nonlinear switched systems

# A basic Example

An example of controlled switched systems

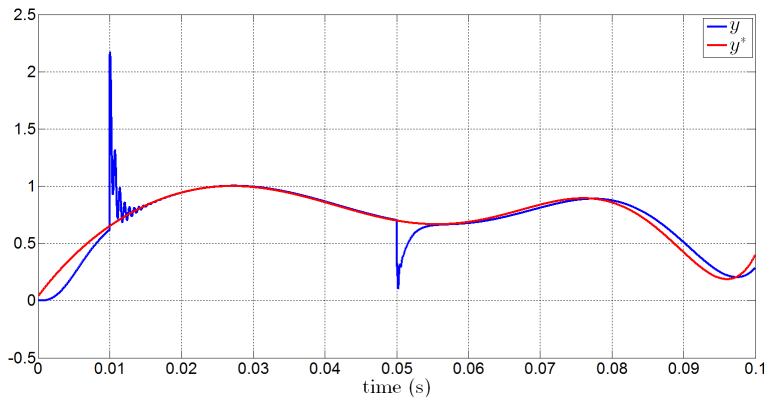
- Consider controlling three linear "unknown" systems that switch



$$\Sigma_p(u \mapsto y) := \begin{cases} \dot{x}(t) = A_p x(t) + B_p u(t) \\ y = C_p x(t) \end{cases}$$

# A basic Example

An example of controlled switched systems



- Switch of the model at  $t_1 = 0.01$  s and  $t_2 = 0.05$  s



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# Para-model control of the HIV-1 model

Presentation

**(Craig, Xia, Venter - 2004)**

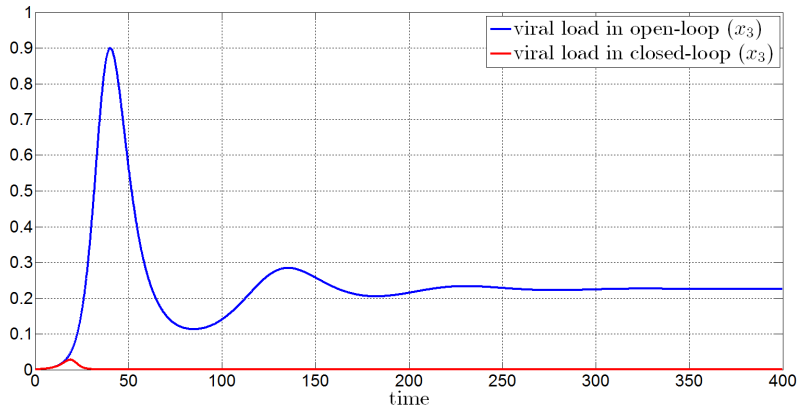
- Control of the predator-prey like model that describes the evolution of the HIV-1 dynamics when subjected to an external "medical agent"
- We do not take into account the constraints that are medically imposed

$$\begin{cases} \dot{x}_1 = s - dx_1 - (1 - u_1)\beta x_1 x_3 \\ \dot{x}_2 = (1 - u_1)\beta x_1 x_3 - \mu x_2 \\ \dot{x}_3 = (1 - u_2)kx_2 - cx_3 \\ y = \begin{pmatrix} 0 & 0 & \gamma \end{pmatrix} x \end{cases}$$

# Para-model control of the HIV-1 model

## Control of the viral load

- Open-loop vs closed-loop



# Para-model control of the HIV-1 model

## Control of the viral load

- the associated  $u$  variable

