# Deep Reinforcement Learning for Bayes-Adaptive Impulse Control of PDMPs

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### Medical context

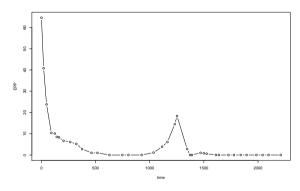


FIGURE: Example of patient data<sup>a</sup>

- Patients who have had cancer benefit from regular follow-up;
- The concentration of clonal immunoglobulin is measured over time;
- The doctor has to make new decisions at each visit.

<sup>&</sup>lt;sup>a</sup>IUCT Oncopole and CRCT, Toulouse, France

### Medical context

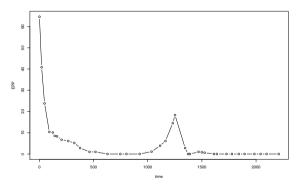


FIGURE: Example of patient data<sup>a</sup>

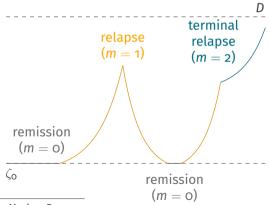
- The concentration of clonal immunoglobulin is measured over time;
- The doctor has to make new decisions at each visit.
- ⇒ Optimising decision-making to ensure the patient's quality of life

Patients who have had cancer benefit from regular follow-up;

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# Controlled PDMP<sup>1</sup>

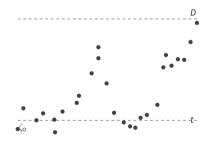
We switch randomly from one deterministic regime to another.



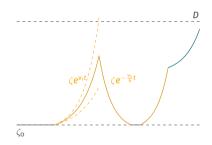
<sup>&</sup>lt;sup>1</sup>Piecewise Deterministic Markov Processes

# Difficulties

#### **Partial observation**

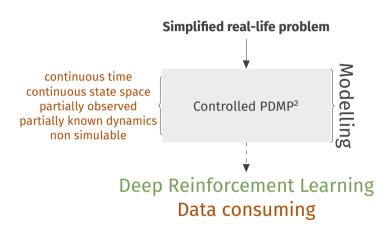


## Partially known dynamics



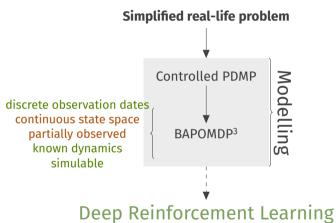
Hypothesis:  ${\bf v_1} \sim {\bf Log\text{-Normal}} \ (\mu, \sigma^{-2})$ , with  $\mu$  and  $\sigma$  unknown.

### Methods



<sup>&</sup>lt;sup>2</sup>Piecewise Deterministic Markov Processes

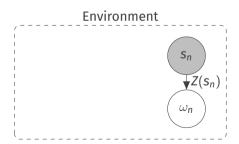
## Methods



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<sup>&</sup>lt;sup>3</sup>Bayes-Adaptive Partially Observed Markov Decision Process



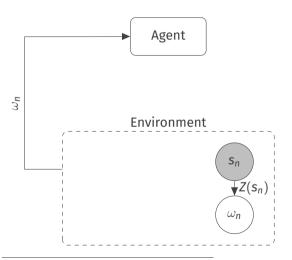


#### POMDP DEFINITION

A POMDP is defined by a tuple ( $\mathbb{S}$ ,  $\mathbb{A}$ , P,  $\Omega$ , Z, c).

- Patient condition  $s = (m, k, \zeta, u) \in S$ ;
- Actions  $a = (\ell, r) \in \mathbb{A}$ ;
- Transition function P(s'|s, a);
- Observation  $\omega = (k, \mathit{F}(\zeta, \epsilon), \mathbb{1}_{m=3}) \in \Omega$ ;
- Observation function  $Z(\omega|s)$ ;
- Cost function  $c : \mathbb{S} \times \mathbb{A} \times \mathbb{S} \to \mathbb{R}$ .

<sup>&</sup>lt;sup>4</sup>Partially Observed Markov Decision Process

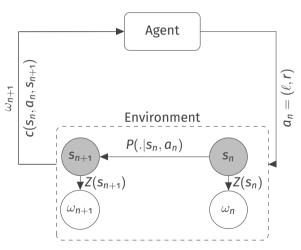


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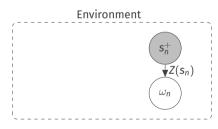
The transition function P(s'|s,a) is a combination of PDMP local characteristics.

<sup>&</sup>lt;sup>4</sup>Partially Observed Markov Decision Process

# Handle uncertainty with Bayesian framework

Normal-Inverse-Gamma( $\Theta$ ) prior patients

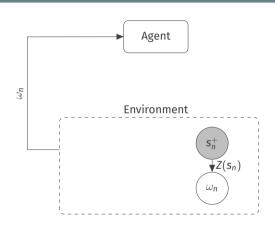




#### BAPOMDP DEFINITION

- Space of hyperstate  $\mathbb{S}^+ = \mathbb{S} \times \Theta$ ;
- Actions  $a = (\ell, r) \in \mathbb{A}$ ;
- Transition function  $P^+(s', \theta'|s, a, \theta)$ ;
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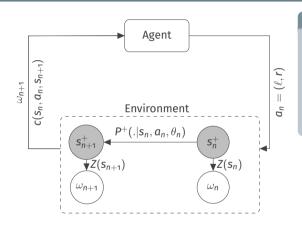
<sup>&</sup>lt;sup>5</sup>Bayes Adaptive Partially observed Markov decision process



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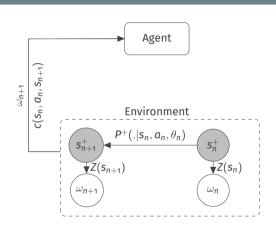


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$$\begin{split} P^+\big((s',\theta') \in B_E \times B_\Theta \mid (s,\theta),a\big) \\ &= \int_{B_\sigma} \mathbf{1}_{B_\Theta} \mathcal{U}(\theta,s,a,s') \times P(ds' \mid s,a,\theta). \end{split}$$

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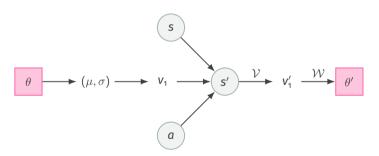
$$P^+((s',\theta')\in B_E\times B_\Theta\mid (s,\theta),a)$$

$$=\int_{B_{\mathsf{F}}} \mathsf{1}_{\mathsf{B}_{\Theta}} \mathcal{U}(\theta,\mathsf{s},a,\mathsf{s}') \times \mathsf{P}(\mathsf{d}\mathsf{s}'\mid \mathsf{s},a,\theta).$$

<sup>&</sup>lt;sup>5</sup>Bayes Adaptive Partially observed Markov decision process

# Generate transition from prior

$$\mathcal{U}(\theta, s, a, s') = \mathcal{W}(\theta, \mathcal{V}(s, a, s')),$$



### Identify an optimal policy $\pi^{\star}$

$$\underbrace{c(s, a, s')}_{\text{Cost function}} = \underbrace{c_V}_{\text{visit cost}} \\ + \underbrace{c_D(H - t') \times \mathbb{1}_{m' = 3}}_{\text{death cost}} \\ + \underbrace{\kappa_C \times r \times \mathbb{1}_{\ell = a}}_{\text{treatment cost}}$$

<sup>&</sup>lt;sup>6</sup>Bayes Adaptative Partially Observable Markov Decision Process

### Identify an optimal policy $\pi^{\star}$

$$\underbrace{V(\pi, \mathbf{S})}_{\text{Optimization criterion}} = \underbrace{\mathbb{E}_{\mathbf{S}}^{\pi} [\sum_{n=0}^{H-1} c(S_{n-1}, A_n, S_n)]}_{\text{Expected total cost as a result of the policy } \pi}$$

<sup>&</sup>lt;sup>6</sup>Bayes Adaptative Partially Observable Markov Decision Process

### Identify an optimal policy $\pi^{\star}$

$$\underbrace{V(\pi,s)}_{\text{Optimization criterion}} = \underbrace{\mathbb{E}_{s}^{\pi} [\sum_{n=0}^{H-1} c(S_{n-1},A_{n},S_{n})]}_{\text{Expected total cost as a result of the policy } \pi}$$

$$\underbrace{V^*(s)}_{\text{Value function}} = \underbrace{\min_{\pi \in \Pi} V(\pi, s)}_{\text{Minimisation across policy space}}$$

<sup>&</sup>lt;sup>6</sup>Baves Adaptative Partially Observable Markov Decision Process

### Identify an optimal policy $\pi^{\star}$

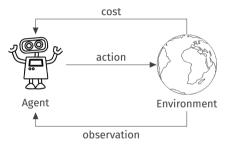
In reality, we do not observe state space!

Let  $h_n = (\omega_0, a_0, \omega_1, a_1, \dots, \omega_n)$  be the history

$$\underbrace{V^{\star}(h)}_{\text{Value function}} = \underbrace{\min_{\pi \in \Pi} V(\pi, h)}_{\text{Minimisation across policy space.}}$$

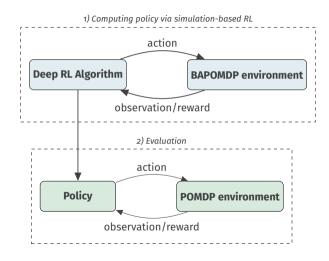
<sup>&</sup>lt;sup>6</sup>Bayes Adaptative Partially Observable Markov Decision Process

# Reinforcement Learning



The optimal policy is obtained from the experiments  $<\omega,a,\omega',c>$ , generated from  $P^+$  transition function

## Evaluate BAPOMDP framework



# Performance BAPOMDP

<b>Policies</b>	Mean Cost (log)	Survival rate	Treatment number	<b>Visit number</b>
ОН	13.45 $\pm$ 0.01	99.60%	0.87	120.57
Random	12.79 $\pm$ 0.01	92.08%	4.36	67.21
Inactive	$8.21 \pm 0.07$	63.42%	0.00	35.39
Threshold	$\textbf{7.03} \pm \textbf{0.04}$	99.98%	5.07	64.67
DQN <sup>7</sup>	$\textbf{5.08} \pm \textbf{0.00}$	99.70%	19.99	38.99
PPO <sup>8</sup>	5.94 $\pm$ 0.01	99.80%	19.99	58.99

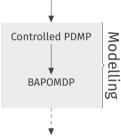
Table: Policy evaluation performance on 5000 Monte-Carlo simulations

<sup>&</sup>lt;sup>7</sup>Deep Q-Network

<sup>&</sup>lt;sup>8</sup>Proximal Policy Optimization

### Conclusion

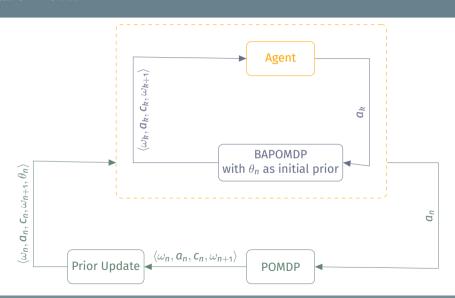
#### Simplified real-life problem



Deep Reinforcement Learning

- Bayes-adaptive method to address the PDMP control problem
- No explicit policies
- No estimates of unknown parameters

# Future work

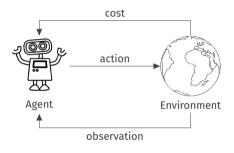


# Policy behavior indicators

Table: Summary of policy behavior indicators based on 5 000 Monte-Carlo simulations.

Indicator	PPO with AM	DQN with AM
Survival rates	99.80 $\%\pm$ 0.00	99.70 $\%$ $\pm$ 0.00
Average number of treatment	19.99 $\pm$ 0.00	19.99 $\pm$ 0.01
Average time spend under treatment	1199.63 $\pm$ 00.04	1199.56 $\pm$ 0.05
Average number of visit	$58.99\pm 0.01$	$\textbf{38.99} \pm \textbf{0.01}$
Average delay between two visits	$\textbf{40.00} \pm \textbf{0.00}$	$\textbf{60.00} \pm \textbf{0.00}$
Rate of visits occurring within 15 days	$\textbf{0.01} \pm \textbf{0.00}$	$\textbf{0.00} \pm \textbf{0.00}$
Rate of visits occurring within 30 days	$66.66\pm 0.17$	$\textbf{0.00} \pm \textbf{0.00}$
Rate of visits occurring within 60 days	$33.33\pm0.17$	$\textbf{100} \pm \textbf{0.00}$

# Reinforcement Learning



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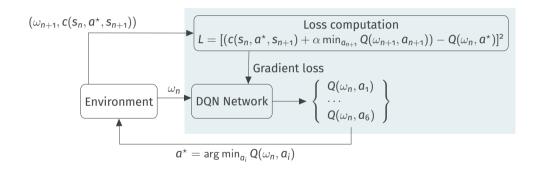
$$\underbrace{Q^{\pi}(s,a)}_{\text{Q value}} = \underbrace{\mathbb{E}^{\pi}[\sum_{n=0}^{H-1}c(S_{n-1},A_n,S_n)|s,a=(\ell,r)]}_{\text{Value of an action in a state according to the policy }\pi$$

$$\underbrace{Q^*(s,a)}_{Q \text{ function}} = \min_{\pi \in \Pi} Q^{\pi}(s,a)$$

$$A(s,a) = Q(s,a) - V(s)$$

Advantage function Extra cost obtained by the agent by taking the action

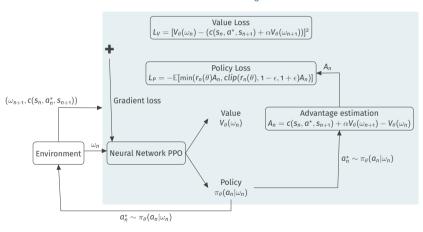
# Algorithm example: DQN<sup>9</sup>



<sup>&</sup>lt;sup>9</sup>Deep Q-Network

# Algorithm example: PPO<sup>10</sup>

#### Agent



<sup>&</sup>lt;sup>10</sup>Proximal policy optimization