

# An example of medical treatment optimization under model uncertainty

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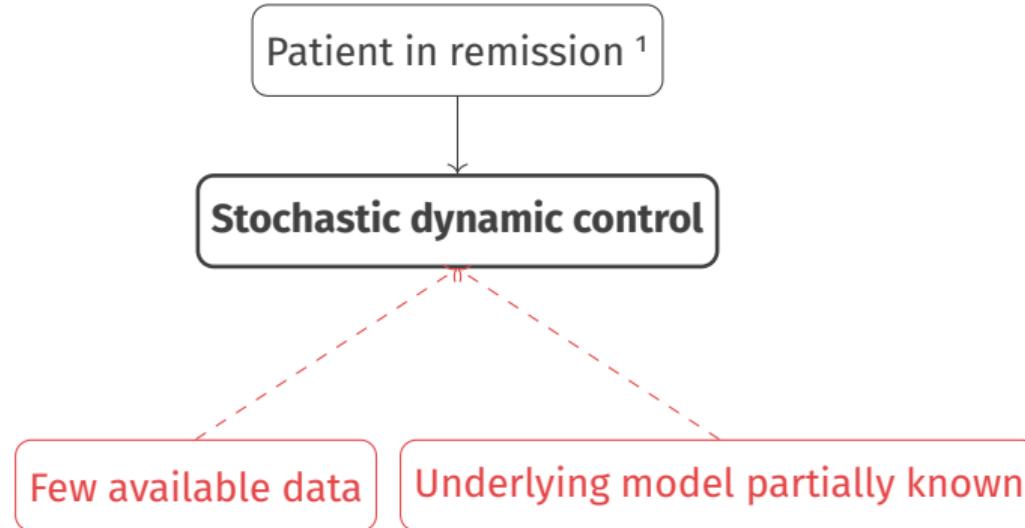
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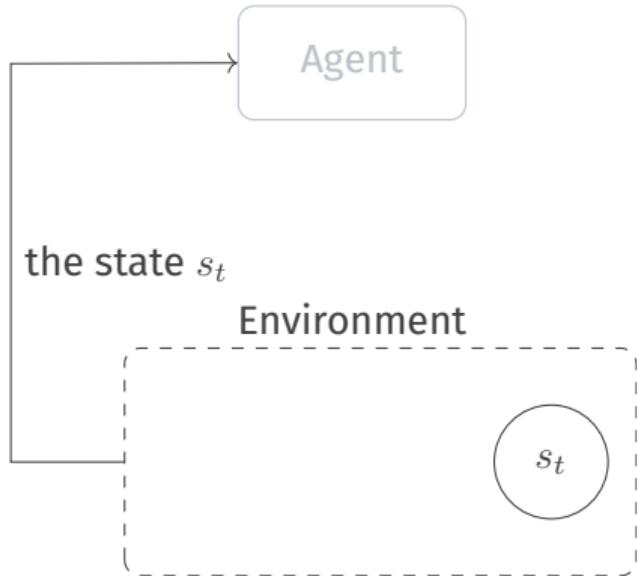
# A medical context



**How can these issues be addressed in a simplified problem ?**

<sup>1</sup>Data from IUC Oncopole, Toulouse, and CRCT, Toulouse, France

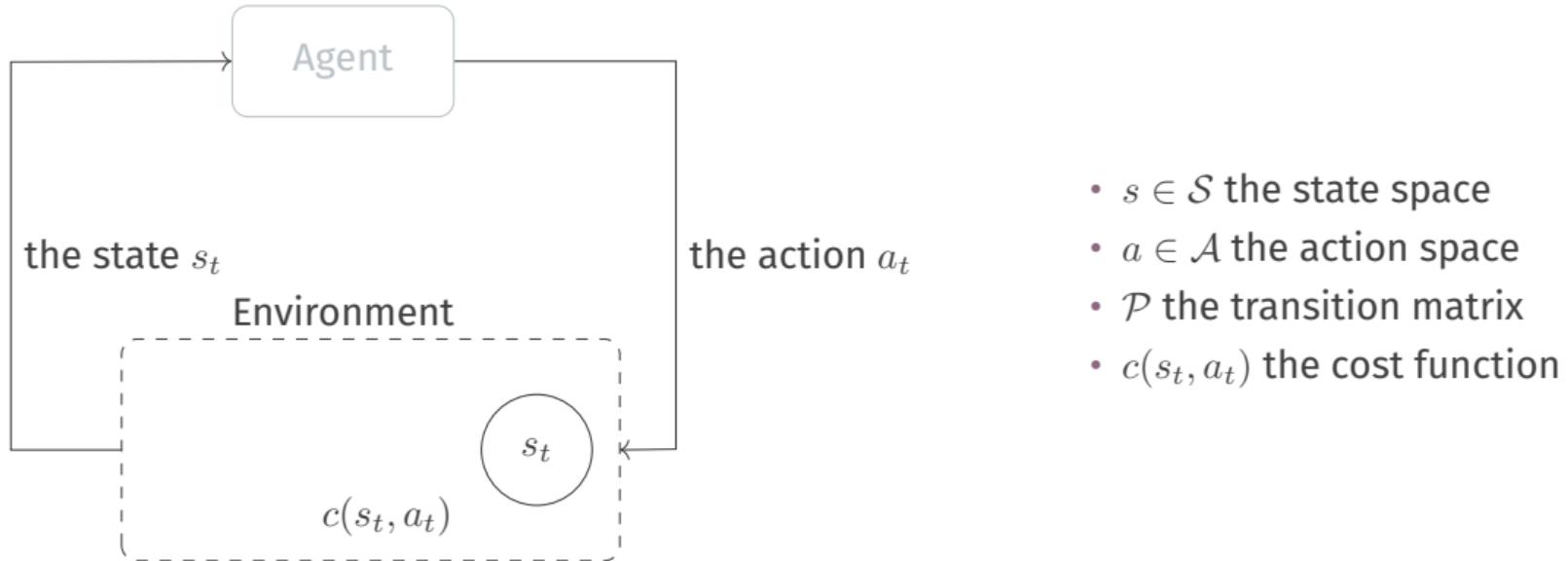
# Markov Decision Process (MDP<sup>2</sup>)



- $s \in \mathcal{S}$  the state space
- $a \in \mathcal{A}$  the action space
- $\mathcal{P}$  the transition matrix
- $c(s_t, a_t)$  the cost function

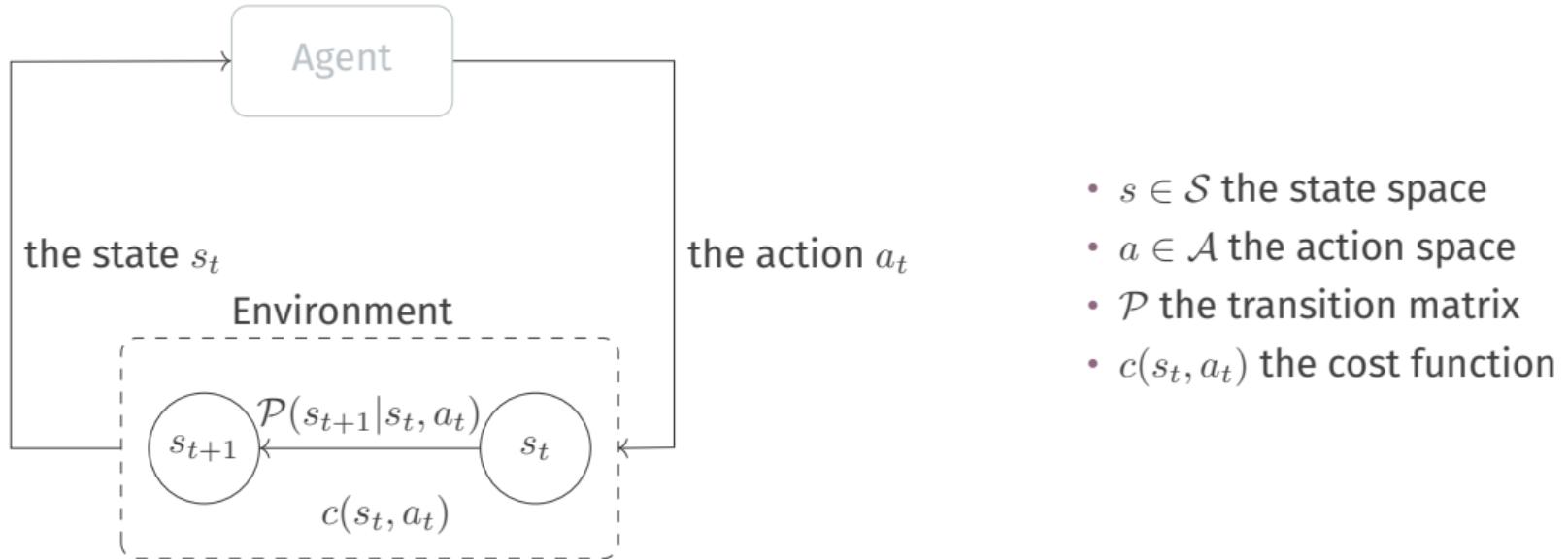
<sup>2</sup>ML Puterman (1994). “Finite-horizon Markov decision processes”. In: Markov Decision Processes: Discrete Stochastic Dynamic Programming. New York: Wiley-Interscience, pp. 78–9.

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# State transition

The transition matrix is partially known

Table: Transition matrix when patient has no treatment ( $a = \emptyset$ ).

$s_t \setminus s_{t+1}$	(0, 0, 0)	(1, 0, 1)	(1, 0, 2)	(1, 1, 1)	(1, 1, 2)	(1, 2, 1)	(1, 2, 2)	(1, 3, 1)	(1, 3, 2)	(2, 4, 0)
(0, 0, 0)	$p_{(0,0,0)}^\emptyset$	$p_{(1,0,1)}^\emptyset$	$p_{(1,0,2)}^\emptyset$	0	0	0	0	0	0	0
(1, 0, 1)	0	0	0	1	0	0	0	0	0	0
(1, 0, 2)	0	0	0	0	0	0	1	0	0	0
(1, 1, 1)	0	0	0	0	0	1	0	0	0	0
(1, 1, 2)	0	0	0	0	0	0	0	0	1	0
(1, 2, 1)	0	0	0	0	0	0	0	1	0	0
(1, 2, 2)	0	0	0	0	0	0	0	0	0	1
(1, 3, 1)	0	0	0	0	0	0	0	0	0	1
(1, 3, 2)	0	0	0	0	0	0	0	0	0	1
(2, 4, 0)	0	0	0	0	0	0	0	0	0	1

# State transition

The transition matrix is partially known

Table: Transition matrix when patient has treatment ( $a = \rho$ ).

$s_t \setminus s_{t+1}$	(0, 0, 0)	(1, 0, 1)	(1, 0, 2)	(1, 1, 1)	(1, 1, 2)	(1, 2, 1)	(1, 2, 2)	(1, 3, 1)	(1, 3, 2)	(2, 4, 0)
(0, 0, 0)	$p_{(0,0,0)}^\rho$	$p_{(1,0,1)}^\rho$	$p_{(1,0,2)}^\rho$	0	0	0	0	0	0	0
(1, 0, 1)	1	0	0	0	0	0	0	0	0	0
(1, 0, 2)	1	0	0	0	0	0	0	0	0	0
(1, 1, 1)	1	0	0	0	0	0	0	0	0	0
(1, 1, 2)	1	0	0	0	0	0	0	0	0	0
(1, 2, 1)	0	0	0	1	0	0	0	0	0	0
(1, 2, 2)	0	0	0	0	1	0	0	0	0	0
(1, 3, 1)	0	0	0	0	0	0	0	1	0	0
(1, 3, 2)	0	0	0	0	0	0	0	0	1	0
(2, 4, 0)	0	0	0	0	0	0	0	0	0	1

# Solving a MDP

Minimizing a cost

## Policy $\pi$

Let  $f : \mathcal{S} \rightarrow \mathcal{A}$  for all  $s \in \mathcal{S}$  is a decision rule.

A sequence of decision rules  $\pi = (f_0, f_1, \dots, f_{H-1})$  is a policy.

The list of costs:

- Treatment: 300
- Disease 1: 200
- Disease 2: 300
- Death: 1000

## Policy cost and value function

$$J_H(\pi, s) = \mathbb{E}\left[\sum_{t=0}^{H-1} c(s_t, a_t) | \pi, s\right]$$

$$V_H(s) = \inf_{\pi \in \Pi} J_H(\pi, s)$$

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## Optimization criterion

$$V^*(s_t) = \min_{a \in \mathcal{A}} [c(s_t, a_t) + \sum_{s_{t+1} \in \mathcal{S}} \mathcal{P}(s_{t+1} | s_t, a_t) V^*(s_{t+1})]$$

# A model-free method

Q-learning<sup>3,4</sup> algorithm



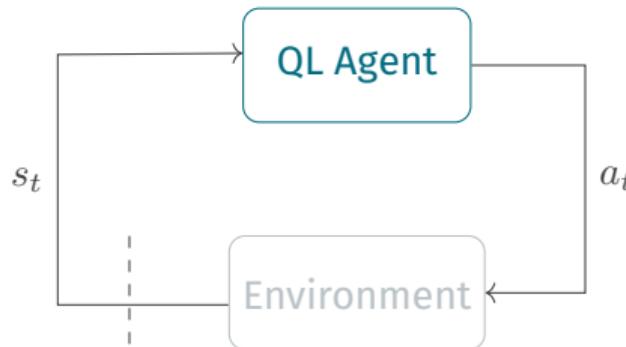
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<sup>3</sup>Christopher J. C. H. Watkins and Peter Dayan (May 1992). “Q-learning”. In: Mach. Learn. 8.3, pp. 279–292.  
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<sup>4</sup>VP Vivek and Dr. Shalabh Bhatnagar (Aug. 2022). “Finite Horizon Q-learning: Stability, Convergence, Simulations and an application on Smart Grids”. In: arXiv:2110.15093v3. DOI: [10.48550/arXiv.2110.15093](https://doi.org/10.48550/arXiv.2110.15093). eprint: [2110.15093v3](https://arxiv.org/abs/2110.15093).

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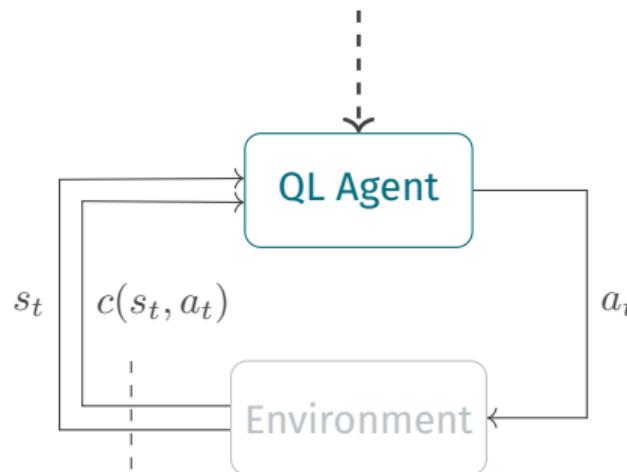
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# A model-free method

Q-learning<sup>3,4</sup> algorithm

$$Q_n(s_t, a_t) = (1 - \alpha)Q_{n-1}(s_t, a_t) + \alpha[c(s_t, a_t) + \min_{a_{t+1} \in \mathcal{A}} Q_{n-1}(s_{t+1}, a_{t+1})]$$



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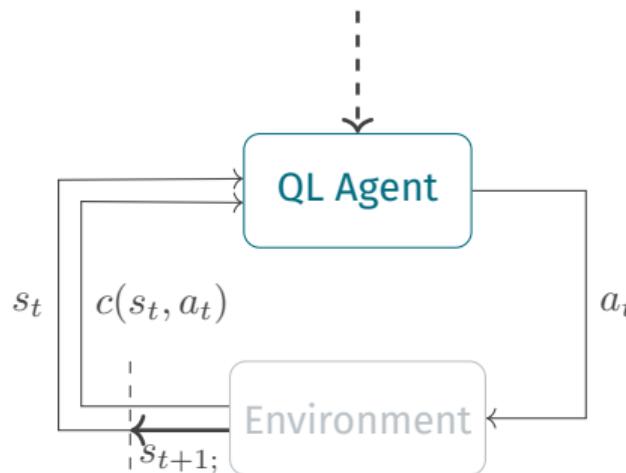
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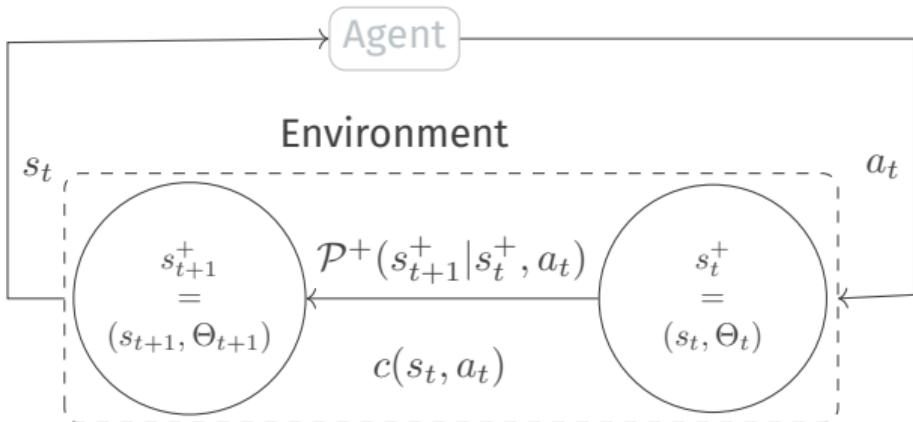
# A bayesian approach

$s_t \setminus s_{t+1}$	(0, 0, 0)	(1, 0, 1)	(1, 0, 2)	(1, 1, 1)	(1, 1, 2)	(1, 2, 1)	(1, 2, 2)	(1, 3, 1)	(1, 3, 2)	(2, 4, 0)
(0, 0, 0)	$p_{(0,0,0)}^\emptyset$	$p_{(1,0,1)}^\emptyset$	$p_{(1,0,2)}^\emptyset$	○	○	○	○	○	○	○

Remark:

- $P(.|s = (0, 0, 0), a = \emptyset) \sim \mathcal{M}(p_{(0,0,0)}^\emptyset, p_{(1,0,1)}^\emptyset, p_{(1,0,2)}^\emptyset)$
- Conjugate distribution :  $f(p^\emptyset | \Theta^\emptyset) \sim \mathcal{D}(\theta_{(0,0,0)}^\emptyset, \theta_{(1,0,1)}^\emptyset, \theta_{(1,0,2)}^\emptyset)$

# Bayes-Adaptive Markov Decision Process (BAMDP<sup>5</sup>)

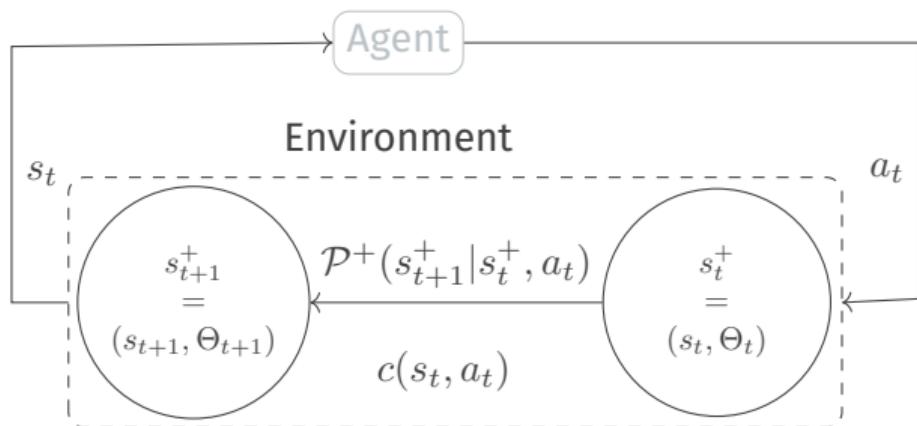


- $s^+ \in \mathcal{S}^+$  the hyper-state space
- $\mathcal{P}^+$  the transition matrix
- $\Theta_{t+1} = \Theta_t + \Delta_{s_{t+1}}^{a_t}$ , with

$$\Delta_{s_{t+1}}^{a_t} = \begin{cases} 1 & \text{if } (s = (0, 0, 0), a_t, s_{t+1}), \\ 0 & \text{else.} \end{cases}$$

<sup>5</sup>Michael O'Gordon Duff (2002). "Optimal learning: Computational procedures for Bayes -adaptive Markov decision processes". PhD thesis. University of Massachusetts Amherst.

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## Optimization criterion

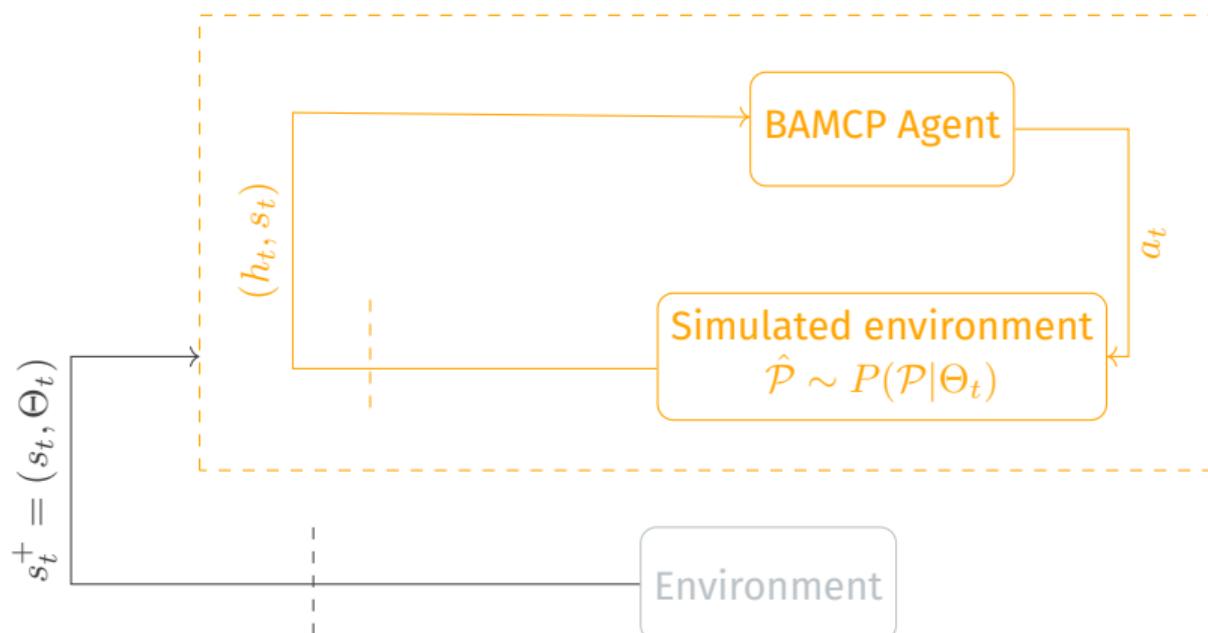
$$V^*(s_t, \Theta_t) = \min_{a \in \mathcal{A}} [c(s_t, a_t) + \sum_{s_{t+1}^+ \in \mathcal{S}^+} \mathcal{P}^+(s_{t+1}^+|s_t^+, a_t) V^*(s_{t+1}, \Theta_{t+1})]$$

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# A model-based method

Bayes-Adaptive Monte-Carlo Planning (BAMCP<sup>6</sup>)

with  $h_t = (s_0, a_0, s_1, \dots, s_{t-1}, a_{t-1})$ ,

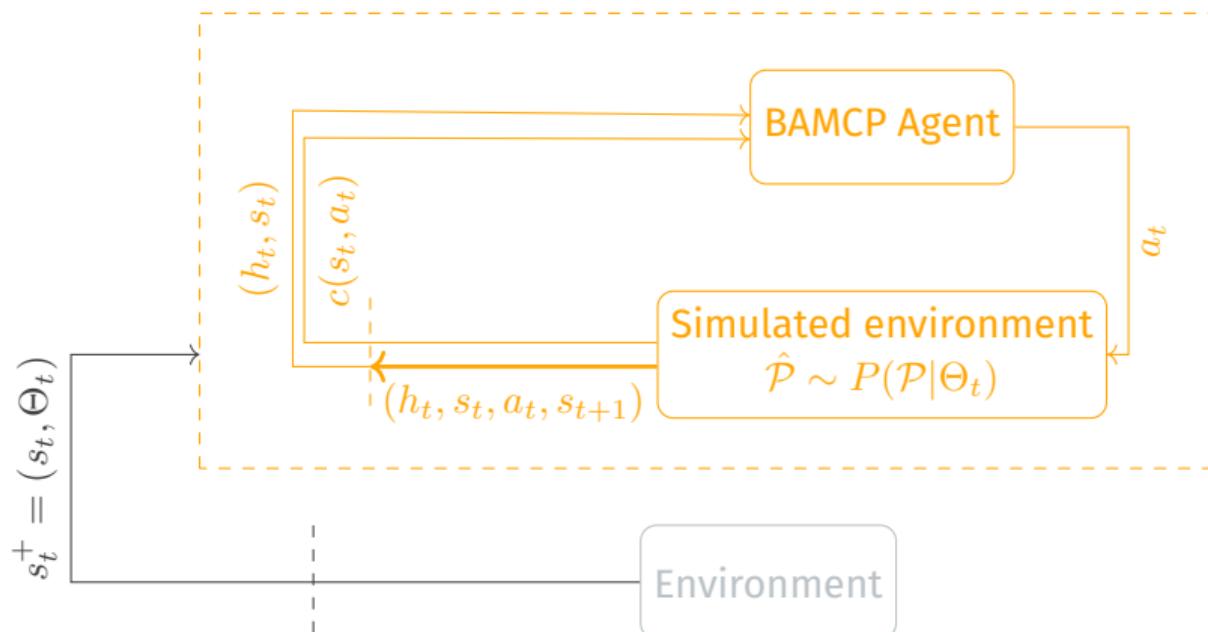


<sup>6</sup>Arthur Guez, David Silver, and Peter Dayan (2012). "Efficient Bayes-Adaptive Reinforcement Learning using Sample-Based Search". In: Advances in Neural Information Processing Systems. Ed. by F. Pereira et al. Vol. 25. Curran Associates, Inc.

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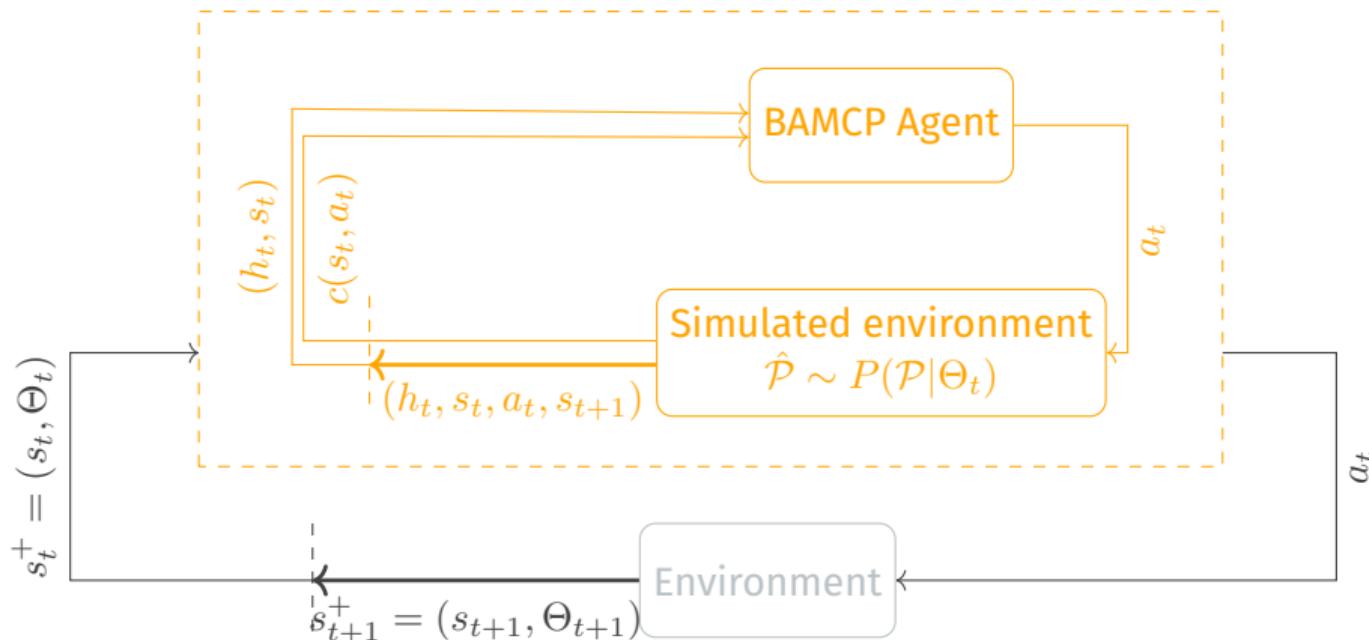


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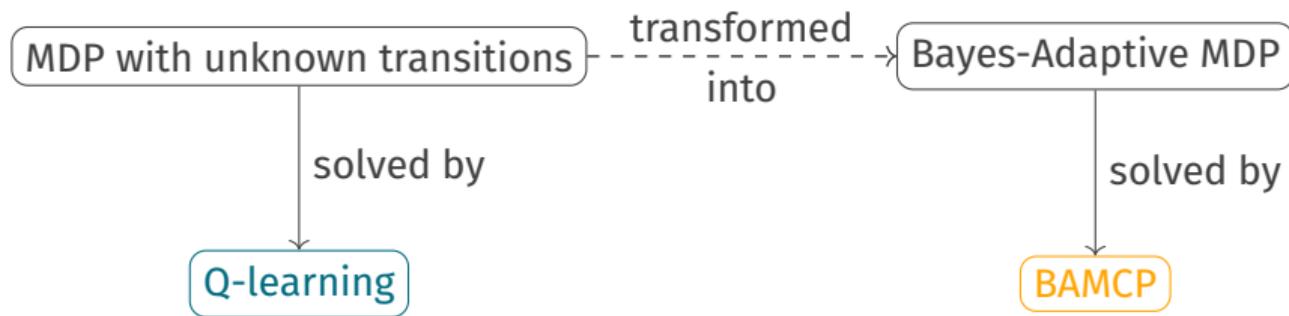
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# Results

The optimal policy exact cost: 888.89

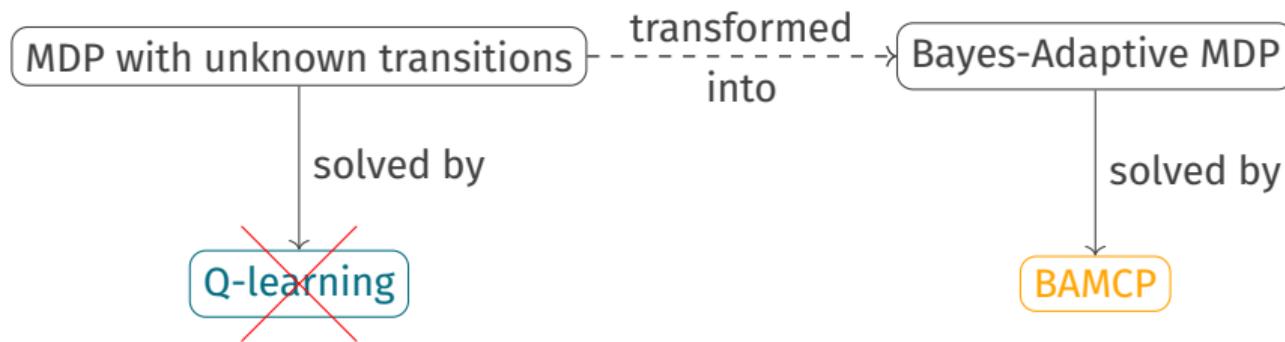
Simulated patients	Q-learning		BAMCP	
	Cost	Time	Cost	Time
$10^2$	$1427.06 \pm 1.05$	0.07 sec	$1302.58 \pm 1.32$	2.07 hours
$10^3$	$936.96 \pm 0.70$	2.48 min	$1297.64 \pm 1.32$	2.22 hours
$10^4$	$936.93 \pm 0.70$	4.17 min	NC	4 days
$10^6$	$891.6 \pm 0.68$	10.21 min	NC	1.5 years

# Conclusion



- ◆ Mathematical framework
- ◆ Model-free method
- ◆ Model-based method

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- ◆ Mathematical framework
- ◆ Model-free method
- ◆ Model-based method

# Perspectives

MDP model	→ PDMP <sup>7</sup> model
Finite state space	→ Continuous state space
Markovian	→ Semi-Markovian
Complete observations	→ Hidden observations

Unlike model-free methods and deep reinforcement learning, **bayesian approaches** do not require as much interaction with the environment.

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<sup>7</sup>Mark H. A. Davis (1984). “Piecewise-Deterministic Markov Processes: A General Class of Non-Diffusion Stochastic Models”. In: *Journal of the Royal Statistical Society Series B (Methodological)* 46, pp. 353–376.