An example of medical treatment optimization under model uncertainty

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June 7, 2023

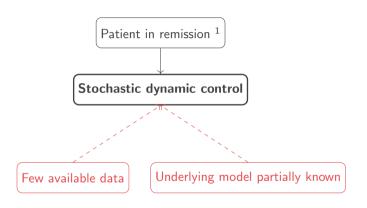






A medical context



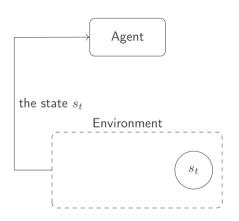


How can these issues be addressed in a simplified problem?

¹Data from IUC Oncopole, Toulouse, and CRCT, Toulouse, France

Markov Decision Process (MDP)

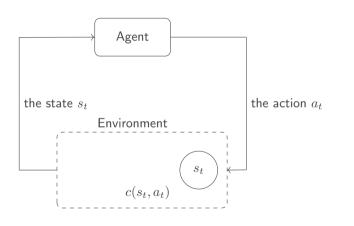




- $s \in \mathcal{S}$ the state space
- ullet $a\in\mathcal{A}$ the action space
- ullet ${\cal P}$ the transition matrix
- $c(s_t, a_t)$ the cost function

Markov Decision Process (MDP)

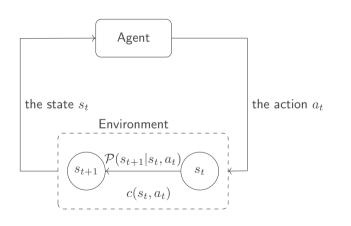




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The transition matrix is partially known



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

| $s_t \backslash s_{t+1}$ | (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 |



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) | $p^{ ho}_{(0,0,0)}$ | $p^{ ho}_{({	extbf{1}},{	extbf{0}},1)}$ | $p^{ ho}_{(1,0,2)}$ | 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 |



The list of costs:

• Treatment: 300

• Disease 1: 200

• Disease 2: 300

• Death: 1000

Policy π

Let $f: \mathcal{S} \to \mathcal{A}$ 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.

Policy cost

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



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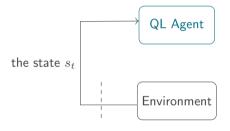
Optimization criterion

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

A model-free method

Q-learning^{2,3} algorithm

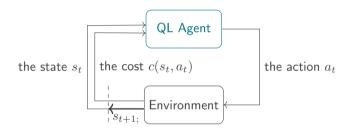




A model-free method

Q-learning^{2,3} algorithm





²Christopher J. C. H. Watkins and Peter Dayan (May 1992). "Q-learning". In: *Mach. Learn.* 8.3, pp. 279–292. ISSN: 1573-0565. DOI: 10.1007/BF00992698.

³VP 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. eprint: 2110.15093v3.

A bayesian approach



Remark:

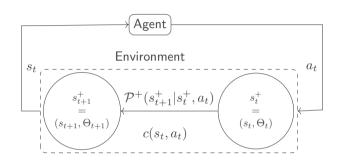
$$\bullet \ \ P(.|s=(0,0,0),a=\emptyset) \sim \mathcal{M}(p_{({\color{blue}0,0,0})}^\emptyset,p_{({\color{blue}1,0,1})}^\emptyset,p_{({\color{blue}1,0,2})}^\emptyset)$$

$$\bullet \ \ \mathsf{Conjugate \ distribution}: \ f(p^\emptyset|\Theta^\emptyset) \sim \mathcal{D}(\theta^\emptyset_{({\color{red}0,0,0})}^\emptyset,\theta^\emptyset_{({\color{red}1,0,1})},\theta^\emptyset_{({\color{red}1,0,2})})$$

•
$$f(p^{\emptyset}|\Theta^{\emptyset}) \propto \prod_{s_{t+1} \in \mathcal{S}} p_{s_{t+1}}^{\emptyset, \theta_{s_{t+1}}^{\emptyset} - 1}$$

Bayes-Adaptive Markov Decision Process (BAMDP4)





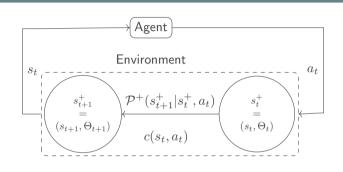
- $s^+ \in \mathcal{S}^+$ the hyper-state space
- ullet \mathcal{P}^+ the transition matrix
- ullet $\Theta_{t+1} = \Theta_t + \Delta^{a_t}_{s_{t+1}}$, with

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

⁴Michael O'Gordon Duff (2002). "Optimal learning: Computational procedures for Bayes -adaptive Markov decision processes". PhD thesis. University of Massachusetts Amherst.

Bayes-Adaptive Markov Decision Process (BAMDP4)





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

Optimization criterion

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

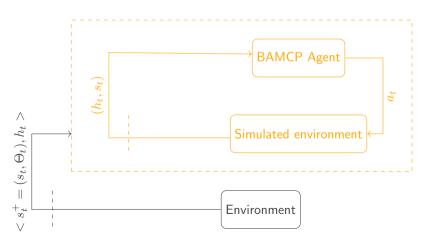
⁴Michael O'Gordon Duff (2002). "Optimal learning: Computational procedures for Bayes -adaptive Markov decision processes". PhD thesis. University of Massachusetts Amherst.

A model-based method

Bayes-Adaptive Monte-Carlo Planning (BAMCP⁵)



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

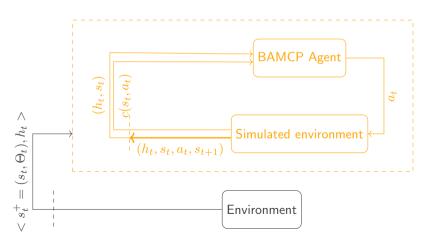


A model-based method

Bayes-Adaptive Monte-Carlo Planning (BAMCP⁵)



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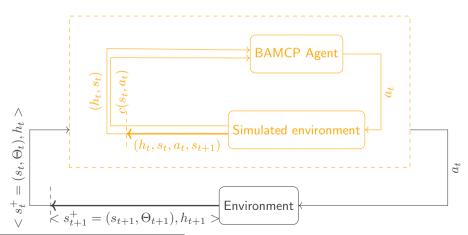


A model-based method

Bayes-Adaptive Monte-Carlo Planning (BAMCP⁵)



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



Results

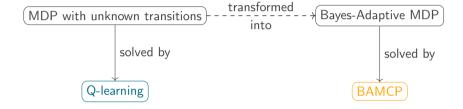


The optimal policy exact cost: 888.89

| Simulated patients | Q-learn | ing | BAMCP | | |
|--------------------|--------------------|-----------|--------------------|-----------|--|
| | Cost | Time | Cost | Time | |
| 10^{2} | 1427.06 ± 1.05 | 0.07 sec | 1377.62 ± 1.21 | 15.41 min | |
| 10^{3} | 936.96 ± 0.70 | 2.48 min | 1340.92 ± 1.04 | 17.86 min | |
| 10^{4} | 936.93 ± 0.70 | 4.17 min | NC | 4 days | |
| 10^{6} | 891.6 ± 0.68 | 10.21 min | NC | 1.5 years | |

Conclusion



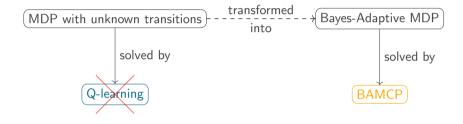


- Mathemathical framework
- Model-free method
- Model-based method

Model-free methods don't work because we don't have enough to interact with.

Conclusion





Mathemathical framework

Model-free method

Model-based method

Model-free methods don't work because we don't have enough to interact with.

Perspectives



 $\mathsf{MDP}\ \mathsf{model} \qquad \qquad \to \ \mathsf{PDMP}^6\ \mathsf{model}$

Finite state space \rightarrow Continuous state space

Markovian ightarrow Semi-Markovian

 ${\sf Complete \ observations} \quad \rightarrow \quad {\sf Hidden \ observations}$

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

⁶Mark 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.