

# Automating anaesthetic delivery with deep reinforcement learning

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# *Automating Anaesthetic Delivery with Deep Reinforcement Learning*

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NEUROSCIENCE  
STATISTICS  
RESEARCH  
LABORATORY



**THE PICOWER  
INSTITUTE**  
FOR LEARNING AND MEMORY

# Outline

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## ***Background***

- Closed loop anaesthetic delivery & reinforcement learning

## ***Methods***

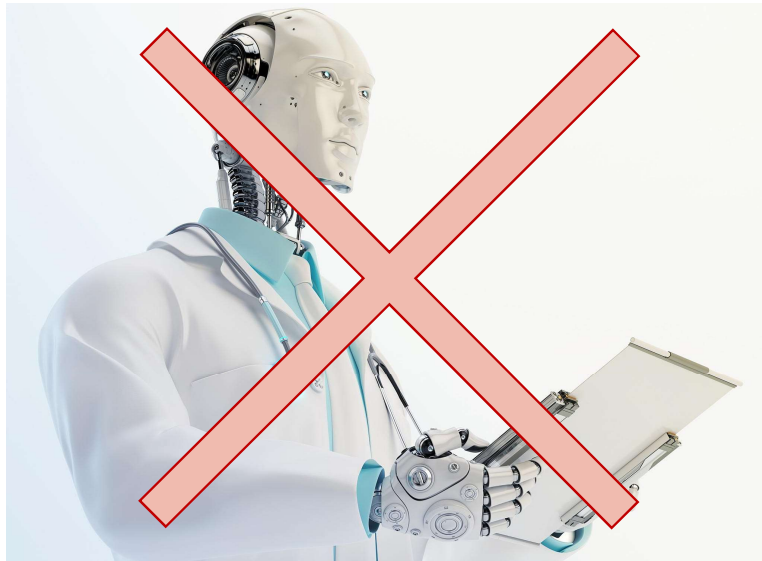
- Deep RL paradigm for controlling level of unconsciousness

## ***Results***

- Simulation study

# Autopilot for Anaesthesia

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# Closed Loop Anesthetic Delivery (CLAD)

Anaesthesiologist sets a *target level of unconsciousness*



Controller determines an appropriate dose



An infusion pump administers the dose to patient

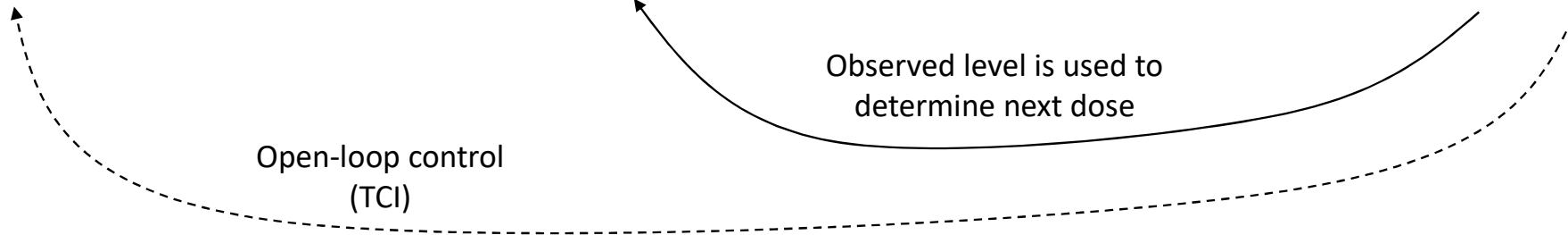


EEG monitor provides *observed level of unconsciousness*



Observed level is used to determine next dose

Open-loop control (TCI)



# CLAD – What's Been Done?

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Paper	Subject	Controller	Level of Unconsciousness
Absalom (2002)	Human	PID	BIS
Dumont (2009)	Simulation	Robust PID/CRONE	WAV <sub>CNS</sub>
Shanechi (2013)	Rodent	LQR	BSP
Moore (2014)	Human	Tabular RL	BIS
<i>Many More...</i>			

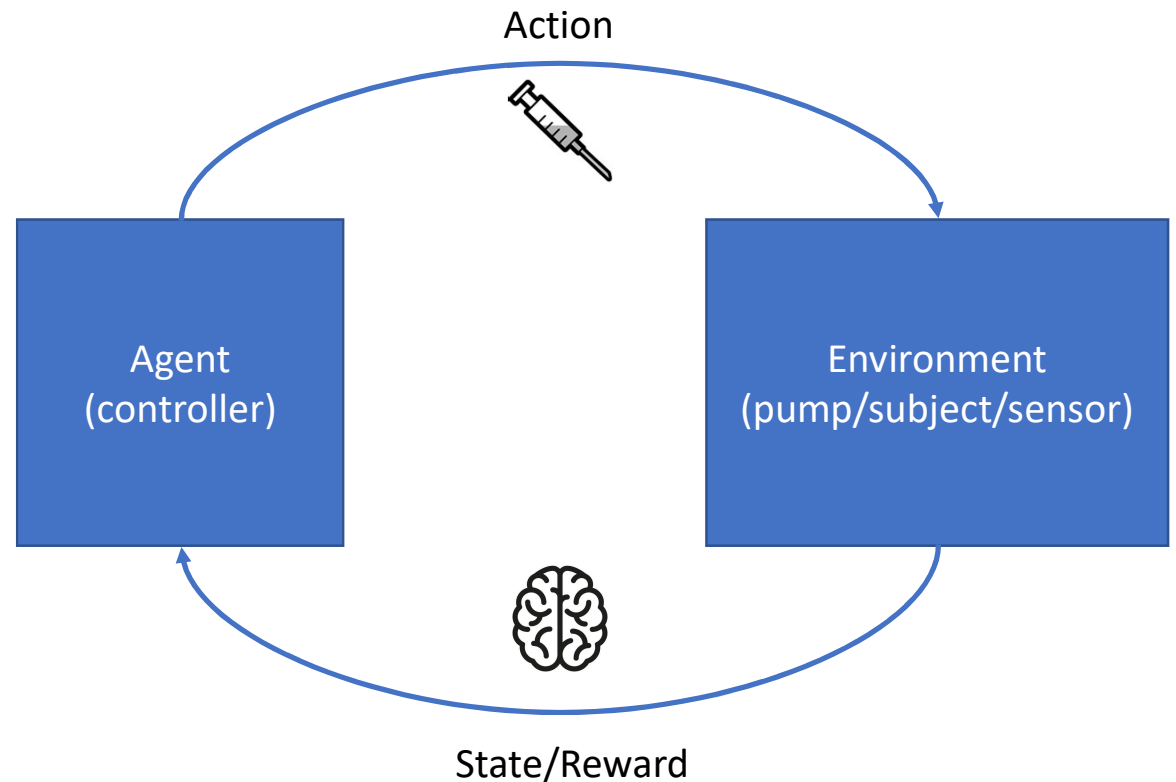
# Reinforcement Learning

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The agent observes a **state**

Based on the state, the agent uses its **policy** to determine an **action**

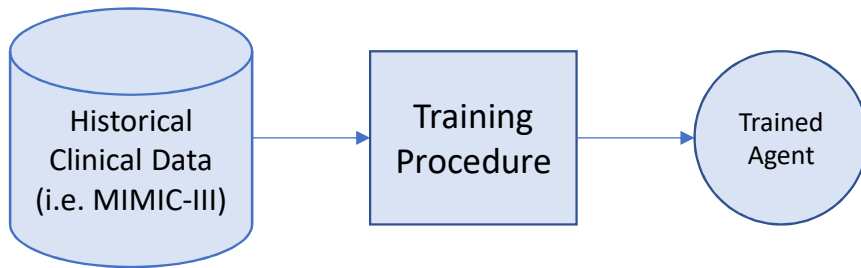
Over time, the agent receives **rewards**, which are used to update its policy



# Reinforcement Learning – Two Approaches

## **Off-Policy:**

*The agent “watches and learns”*



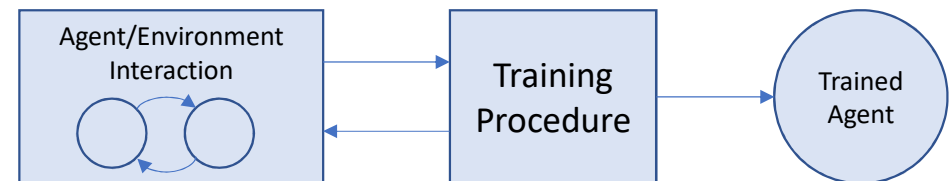
- Agent sees real patient responses
- Agent learns to reflect anaesthesiologist's actions

- Data may not contain enough bad or ideal behavior
- For example, infrequent titration is suboptimal

## **On-Policy:**

*The agent “learns by doing”*

*Focus  
of this  
talk*



- Agent can fully explore action space
- The actions will respond to varying reward functions

- Agent cannot act on humans → requires simulations

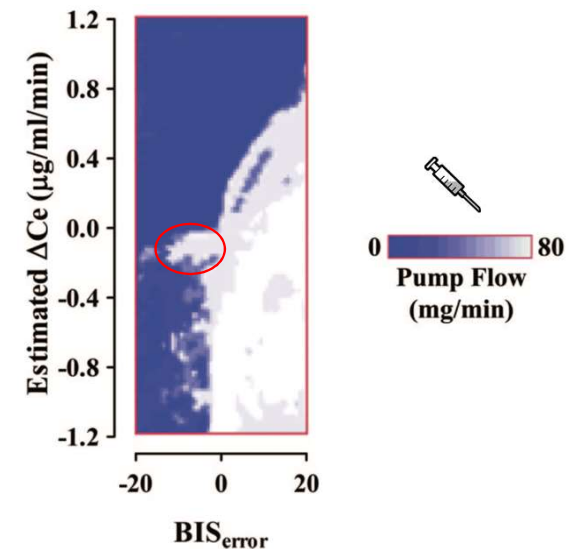


# Tabular RL Policies

A **policy** maps a **state** to an **action**

Tabular RL learns which discrete action should be taken for every *discrete state*

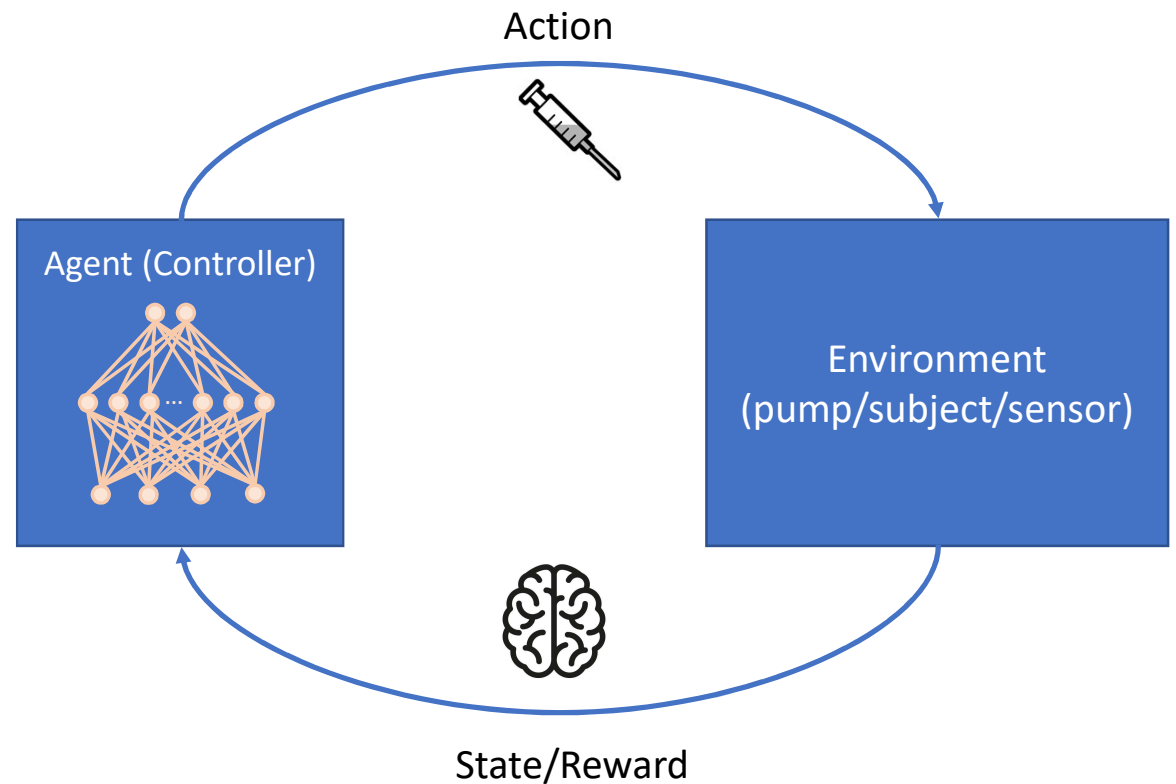
- Ignores the continuous relationship between state and action
- Results in patchy discontinuities
- Dimensionality of the table scales exponentially with the state space



From Moore et al. (2011)

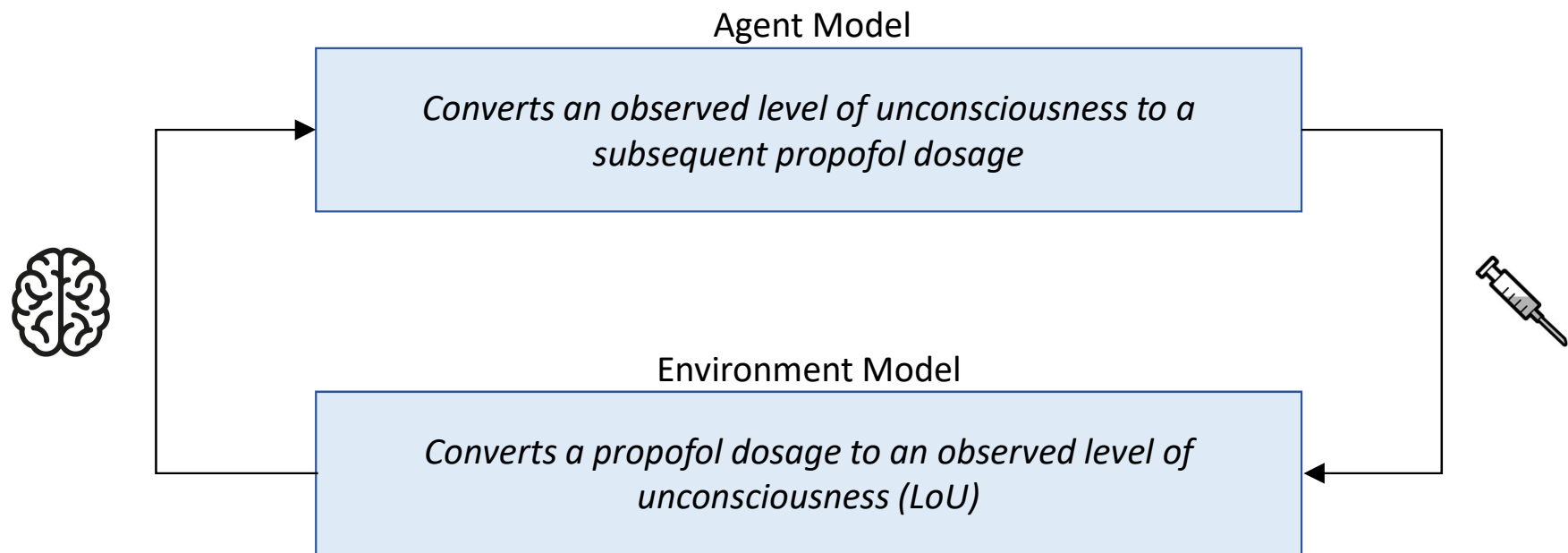
# Deep Reinforcement Learning

The policy is a  
**deep neural network**



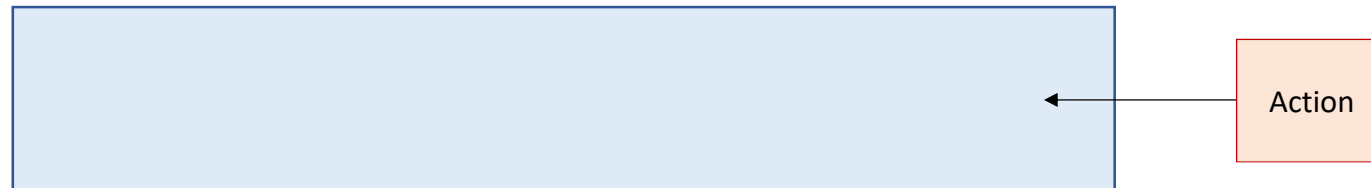
# Deep RL for CLAD

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# The Environment Model

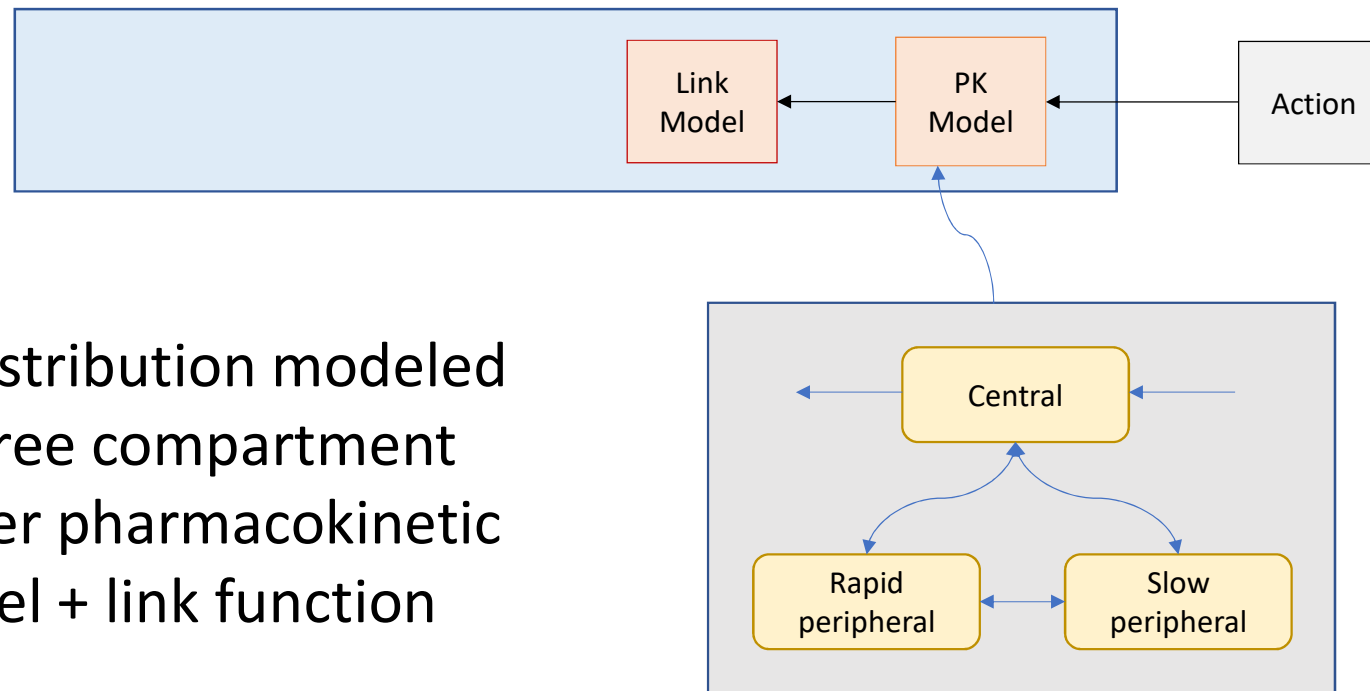
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5 seconds of  
**1.67 mg/s or 0 mg/s**



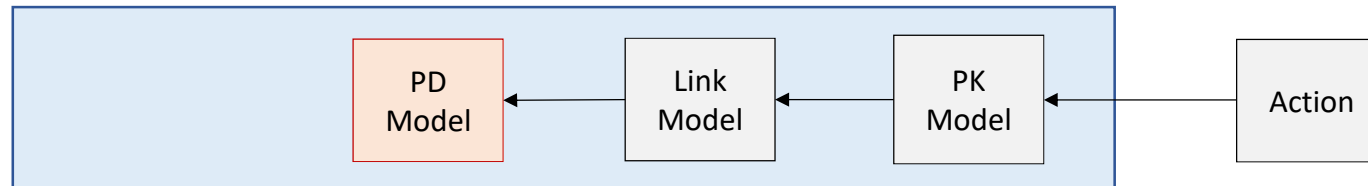
# The Environment Model



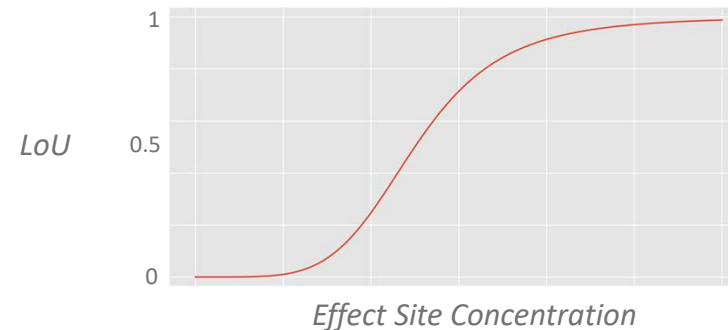
Drug distribution modeled  
by three compartment  
Schnider pharmacokinetic  
model + link function

# The Environment Model

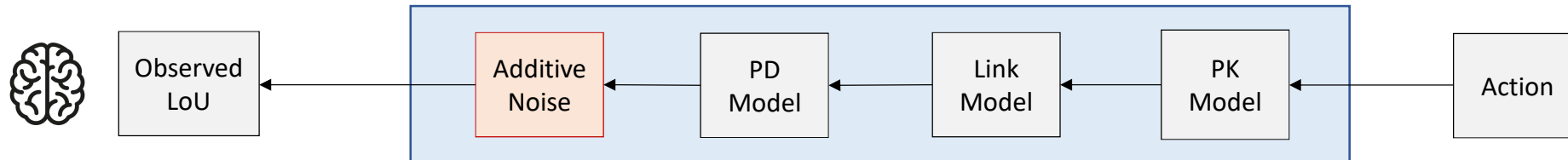
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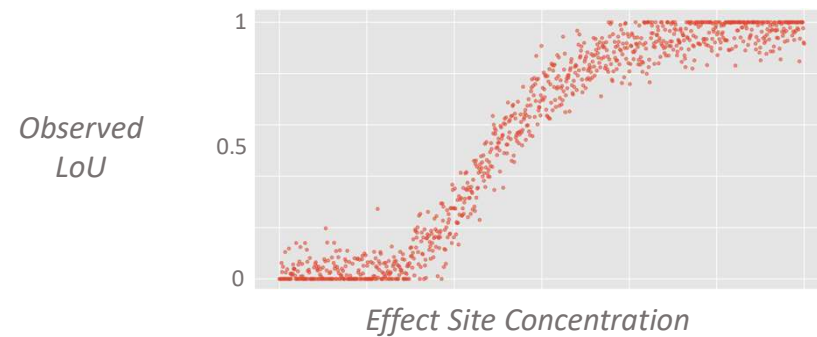
Drug effect modeled by nonlinear pharmacodynamic model determines level of unconsciousness (LoU)



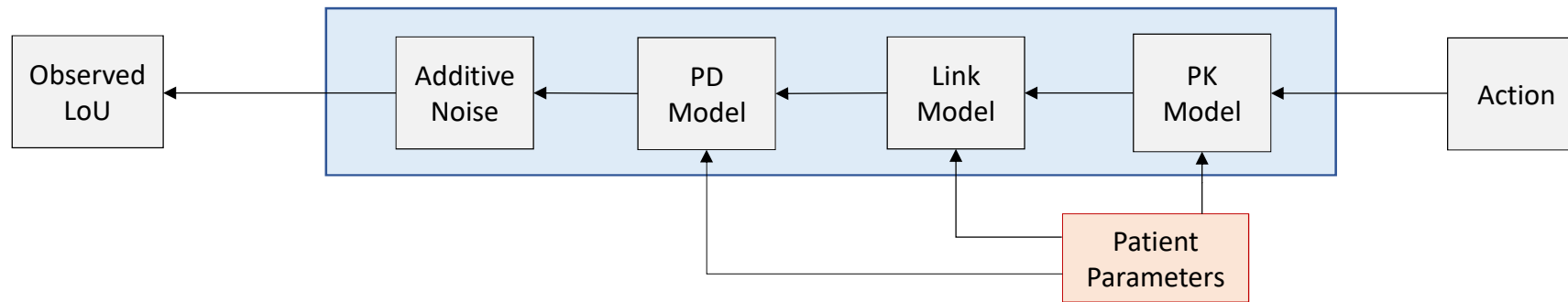
# The Environment Model



Observed LoU includes  
additive noise



# The Environment Model



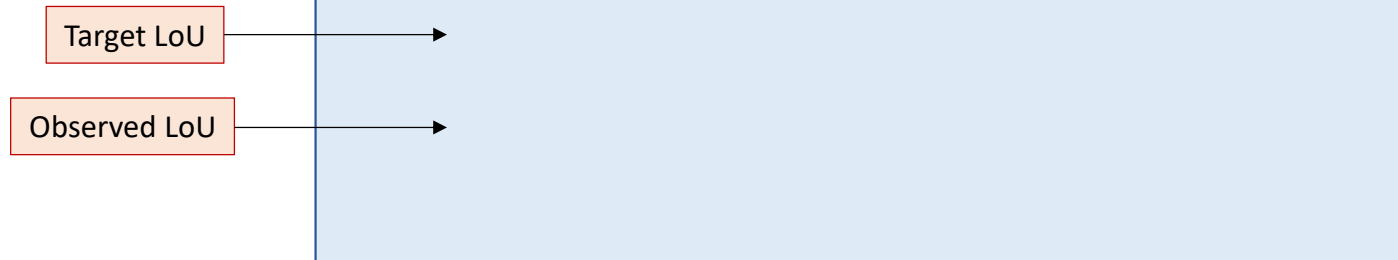
patient variability  
model

Sub-model	Parameter	Units	Generic	Minimum	Maximum
PK	Height	cm	170	160	190
PK	Weight	kg	70	50	100
PK	Age	yr	30	18	90
Link	$k_{e0}$	$\text{min}^{-1}$	0.17	0.128	0.213
PD	$\gamma$	-	5	5	9
PD	$C$	-	2.5	2	6



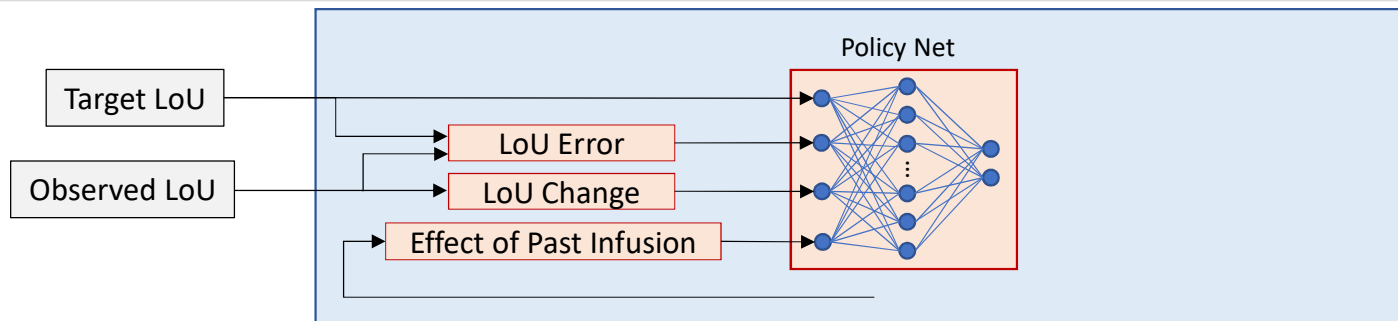
# The Agent Model

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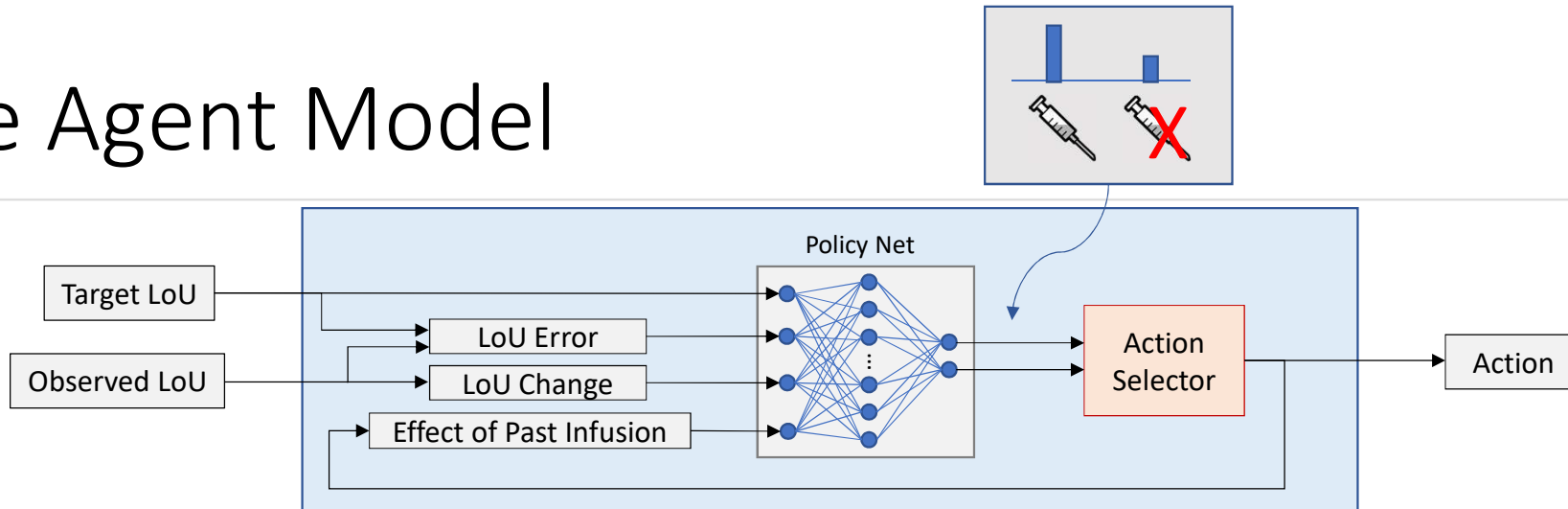
Agent receives observed  
and target LoUs

# The Agent Model



The policy assigns probabilities to actions based on a 4 dimensional observation

# The Agent Model



Agent uses one of three strategies to select an action

## Stochastic

*Picks action randomly according policy output*

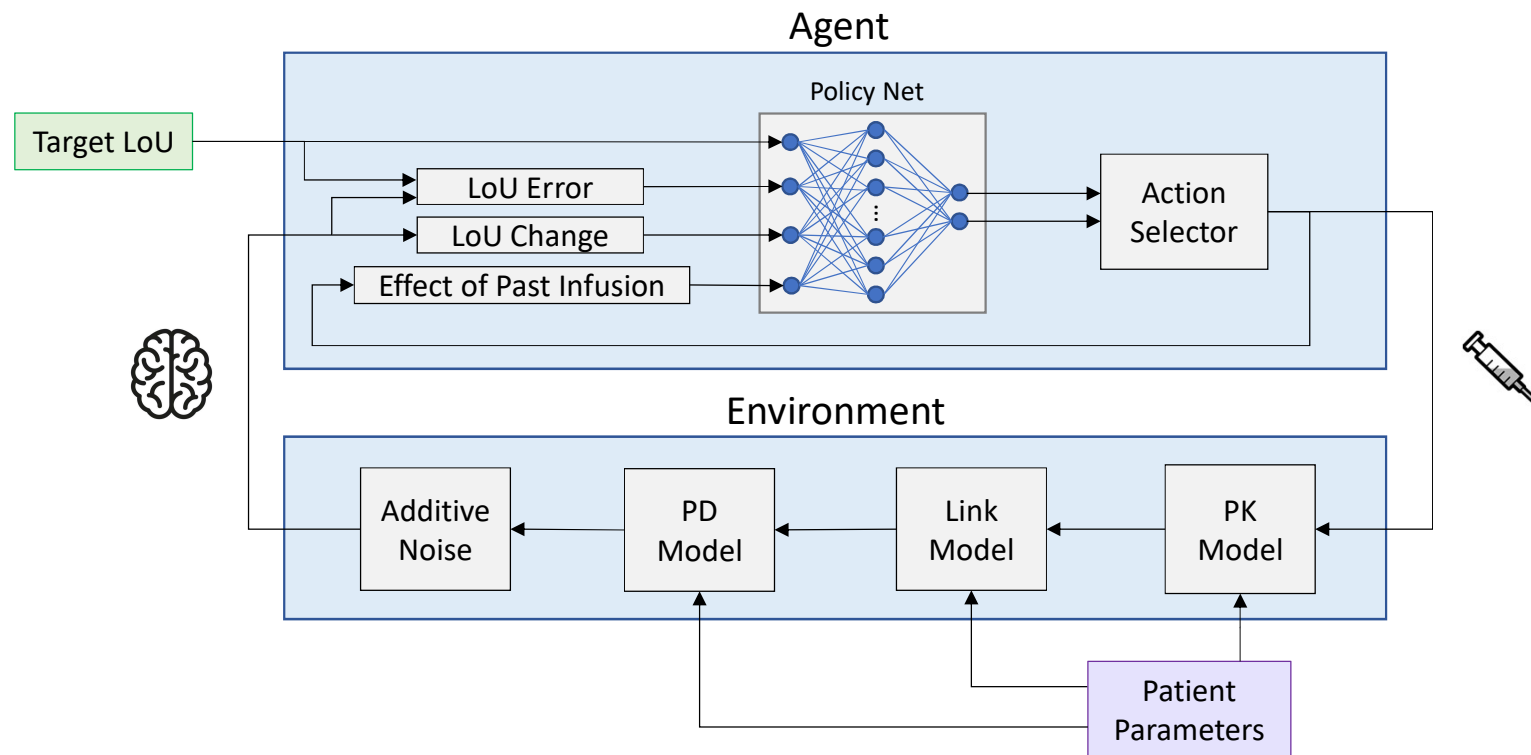
## Deterministic

*Pick action with highest probability*

## Continuous

*Multiply action probability by max dose*

# Complete Simulation Model



# Training the Agent

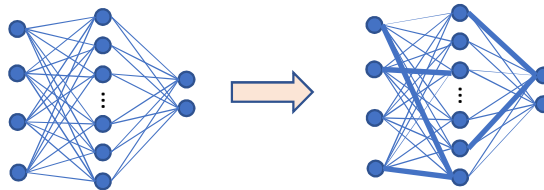
Agent acts with randomness in a series of “episodes”



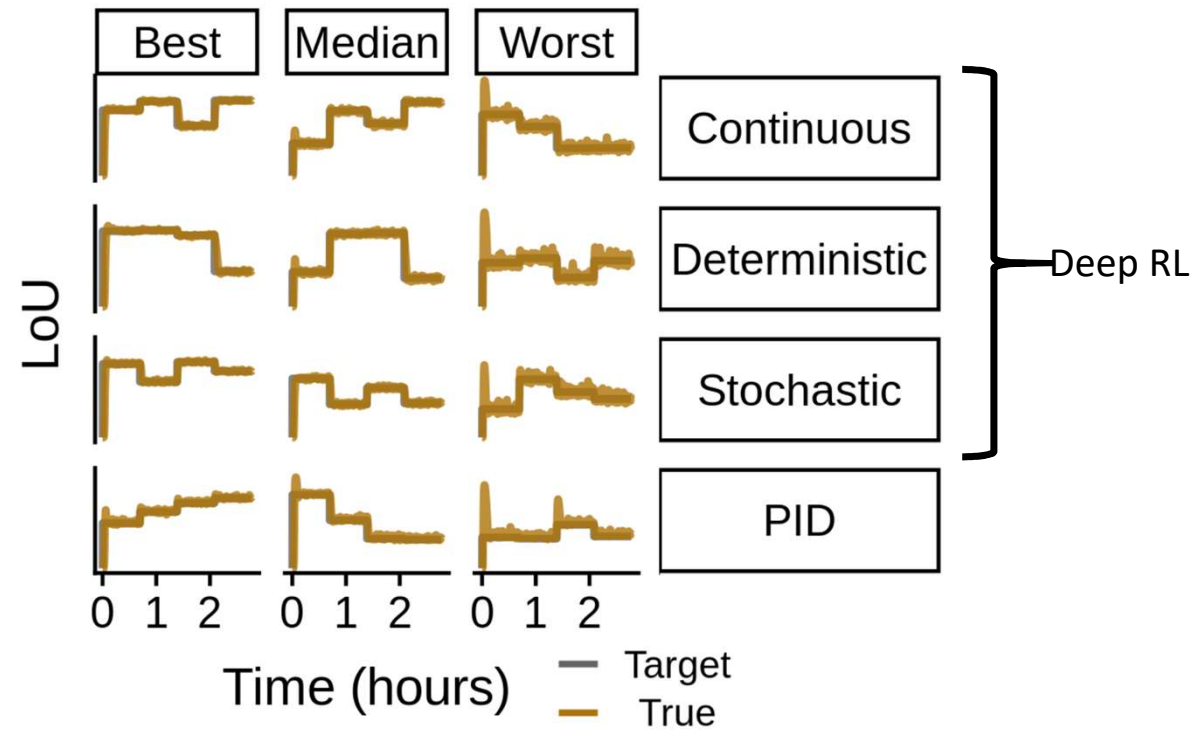
Identify which episodes were best



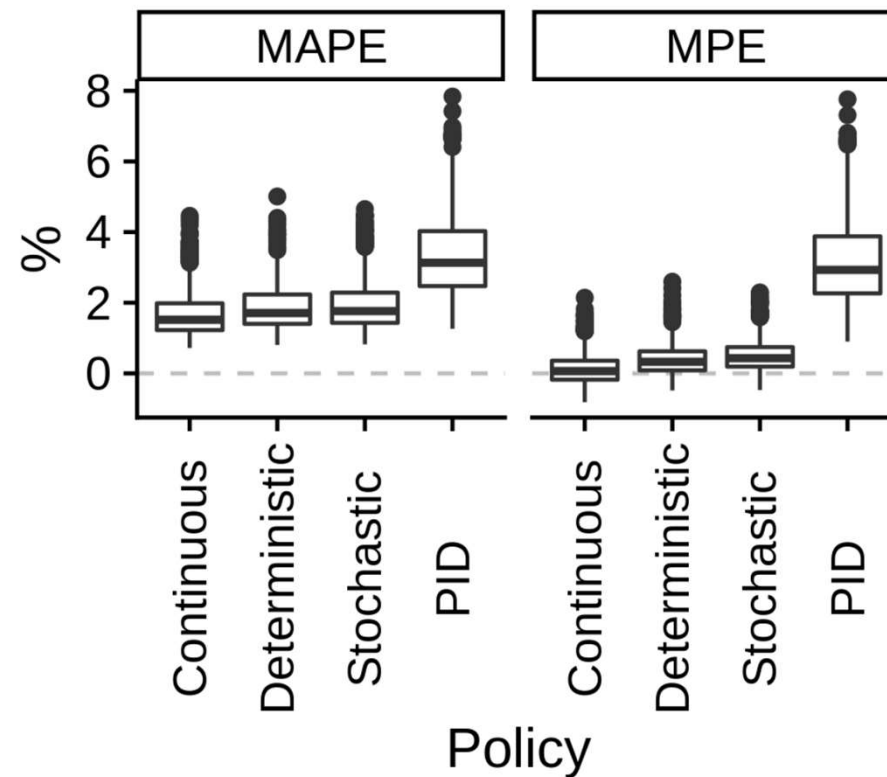
Nudge policy to encourage repeating best actions



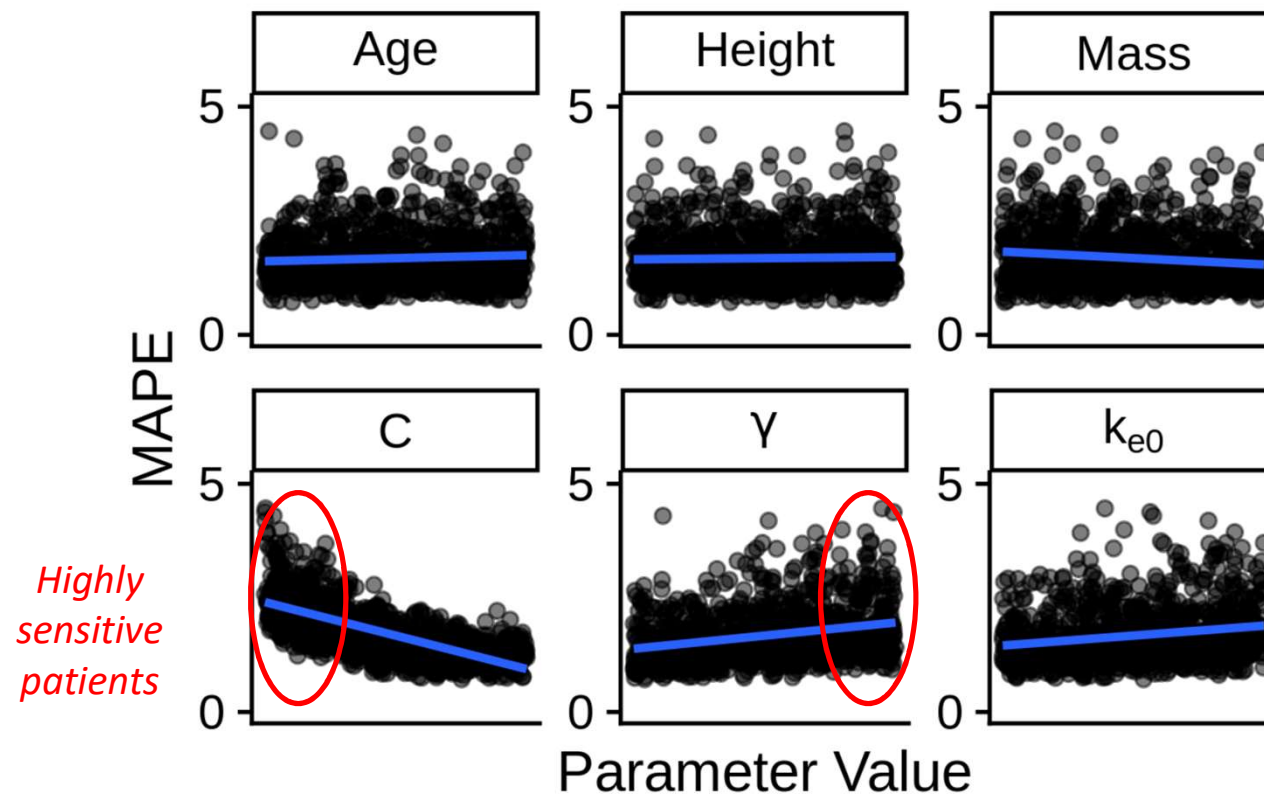
# Deep RL Works



# Deep RL Outperforms PID

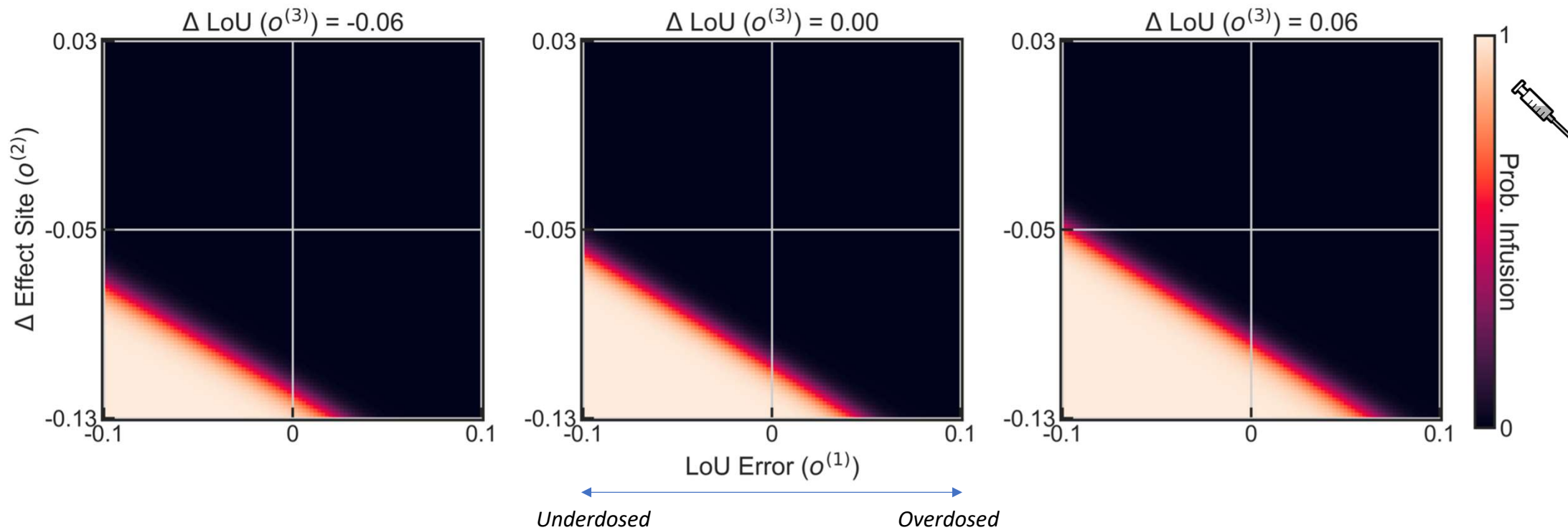


# Deep RL is Robust to Patient Variability





# Deep RL is Not a Black Box



# Key Conclusions

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- Deep RL outperforms PID controller in simulations
- Deep RL resolves issues with tabular RL
  - Enables consideration of other patient variables
  - Reflects continuous state/action relationships
- Resulting policy is not a black box

# Future Work

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- Further simulation studies
  - Intra-patient variability in environment
  - Continued feature engineering
  - Off-policy training on retrospective clinical data
- Clinical studies
  - Human-in-the-loop recommender system
  - Animal studies
  - Volunteers?

# Acknowledgements

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- Marcus Badgeley
- Emery Brown
- Benyamin Meschede-Krasa
- John Abel

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# BACKUP

# A Side by Side Comparison of Control Algorithms

Classical Model-Free (PID)	Classical Model-Based (LQR)	Deep RL
Parameters are <i>tuned</i> using nominal transfer function	Parameters are <i>derived</i> using nominal patient model	Parameters are <i>learned</i> using patient model simulations
Does not optimize	Optimizes a <i>quadratic cost</i>	Optimizes <i>reward</i>
<i>Linear</i> function of <i>error</i>	<i>Linear</i> function of <i>state</i>	<i>Non-Linear</i> function of <i>state</i>
Well-established methods for testing stability, robustness, responsiveness		No formal analysis
End product is a <u><i>deterministic</i></u> controller		