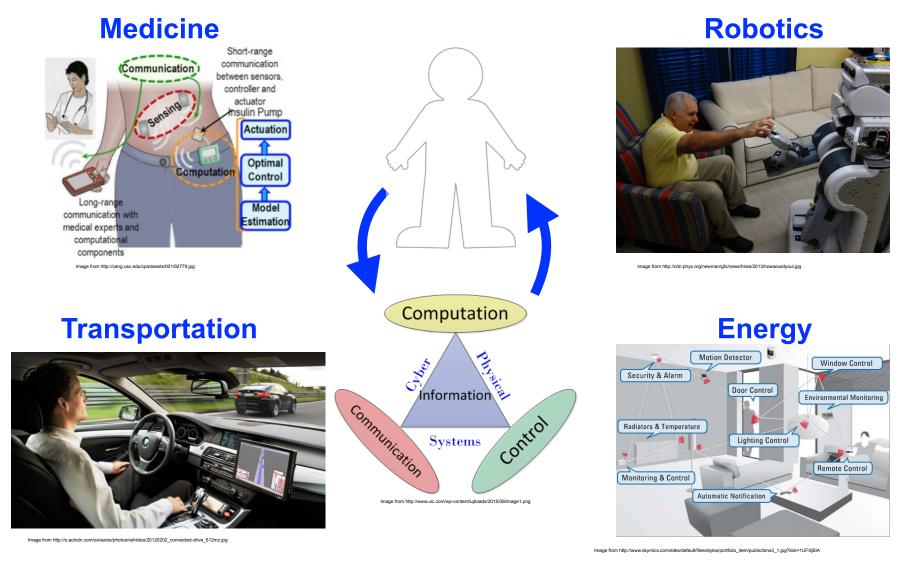
DATA-DRIVEN BEHAVIOR MODELING







Human-in-the-Loop CPS





PRE CISE/19

Challenge: Operator Modeling

- Human-operated CPS
 - CPS operators are integral part of CPS applications
 - Uncertainty due to complex human-automation interaction
 - Needs to be taken into account in controller design and safety analysis
 - Behavior changes over time





Behavioral Modeling in MCPS/IoMT

- Human-in-the-loop MCPS/IoMT
 - Clinicians and/or patients operate and coordinate medical devices
 - Analysis of safety and effectiveness needs to take operator behavior into consideration
 - How much the operator trusts the system
 - When and how operator interferes with automation
- Case study: patient-operated insulin pump
 - Smart pump suggest doses
 - Patients input carb intake
 - Patients can accept or adjust dose
 - How does behavior affect treatment?







A Data-Driven Behavior Modeling and Analysis Framework for Diabetic Patients on Insulin Pumps

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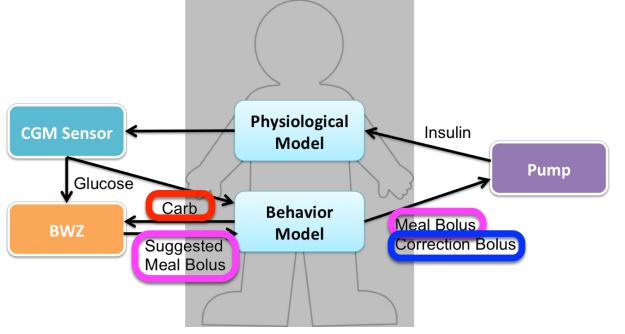


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Type 1 Diabetics (T1D) on Insulin Pumps

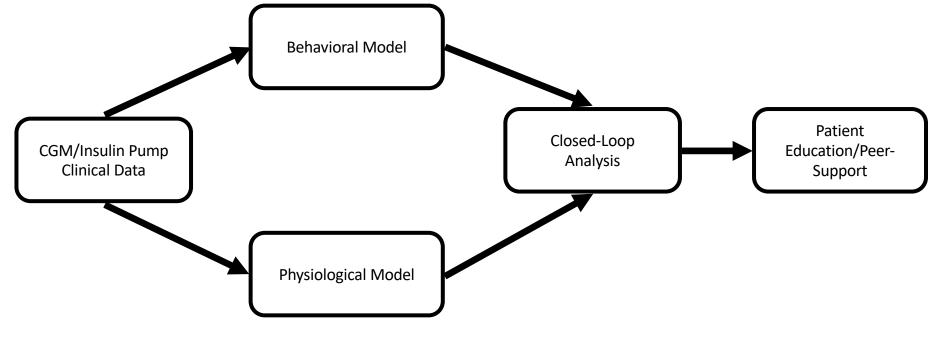
- Sensor-augmented subcutaneous insulin therapy
 - 30% 40% T1D patients in the US use insulin pumps
 - Requires user supervision
 - Input meal information, approve pump-suggested boluses, give non-mealtime boluses, calibrate CGM sensor
 - American Association of Clinical Endocrinologists report highlights critical needs for better understanding the *physiological and psychological* impacts of insulin pumps on diabetic users





Overview of our Approach

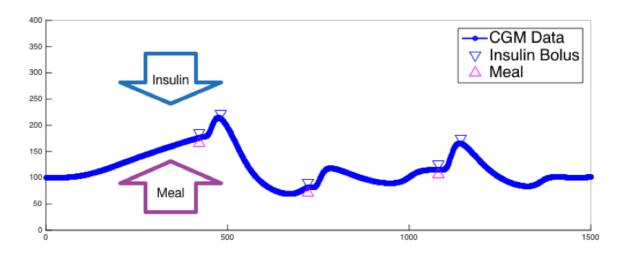
- Behavioral modeling: unsupervised learning
- Physiological modeling: fitting a standard physiological model
- Closed-loop analysis: probabilistic model checking
- Patient education/peer-support: how behaviors affect outcomes





Clinical Dataset

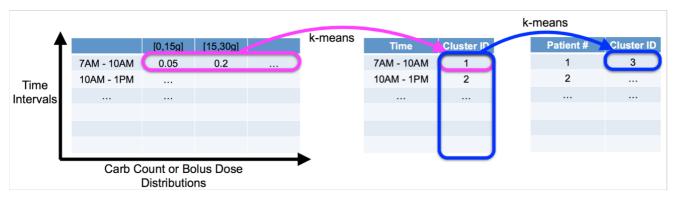
- Sensor-augmented insulin pump data
 - CGM readings, mealtimes & carb counts, pump suggested boluses, userselected boluses
- 55 T1D patients at the UPenn Hospital (age 45.7 ± 15.3, body weight 79.2 ± 21.9 kg)
 - Average time duration 31 days
 - The majority of 84 patients expected to use sensor-augmented insulin pumps
 - 932 T1D patients seen at the UPenn Hospital last year, 60% use insulin pumps, 15% use CGM sensors





Data-Driven Behavior Modeling

- Three behavioral sub-models
 - Eat: distributions of mealtime and carb counts
 - Trust: the likelihood of following pump-suggested boluses and distributions of dose adjustments
 - Correct: distributions of correction-bolus frequencies and doses
- Two-tier clustering heuristic
 - 1st stage: Probability Table —> Vector of Row Cluster ID
 - 2nd stage: Vector of Row Cluster ID —> Patient Cluster ID





Behavioral Modeling in MCPS

Physiological

Model

Behavior

Model

Insulin

Meal Bolus

Correction Bolus

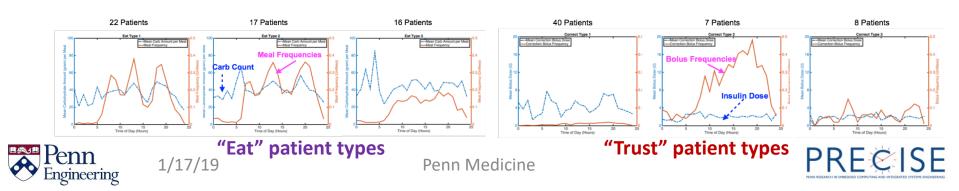
Pump

CGM Senso

Gluco

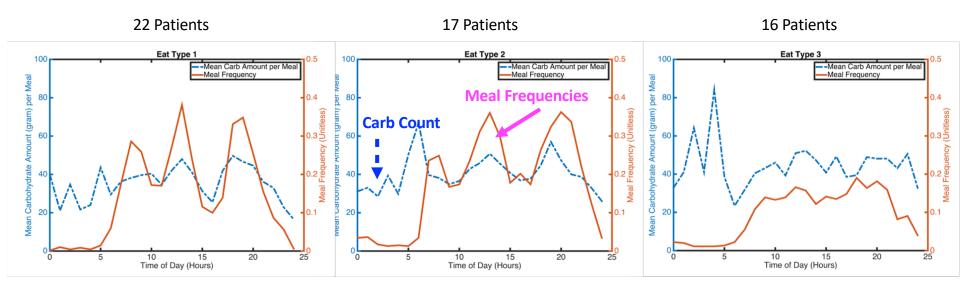
Carb

- "Eat-Trust-Correct" model
 - Distributions of mealtime and carb counts
 - The likelihood of following pump suggestions
 - Distributions of correction frequencies and doses
 - Constructed from actual patient data
 - Identified a set of behavior types
- Combine behavioral and physiological models to assess expected outcomes
 - Immediate implications for patient education



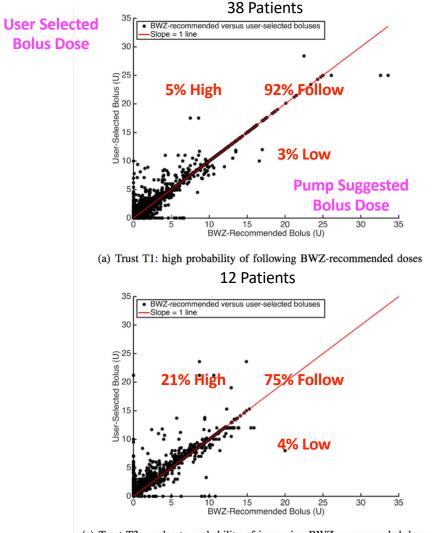
Eat Clusters

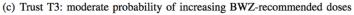
- Eat Type 1: 3 regular meals with low-carb inter-meal snacks
- Eat Type 2: 3 regular meals with moderate-carb inter-meal snacks
- Eat Type 3: no regular meal times

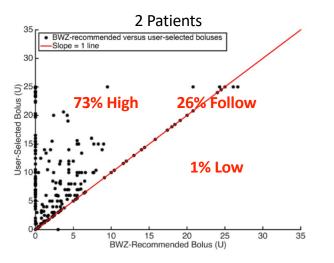




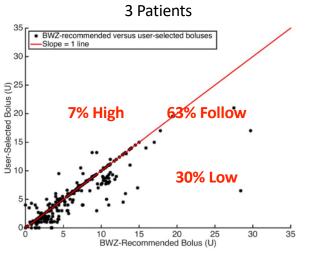
Trust Clusters







(b) Trust T2: high probability of increasing BWZ-recommended doses



(d) Trust T4: high probability of decreasing BWZ-recommended doses



Physiological Model

• Bergman model: compartmental physiological model

$$\begin{array}{c} \text{Plasma Glucose} \longrightarrow \\ \begin{array}{c} G(t) \\ g(t) \\ \\ \frac{d}{dt} \end{array} \\ \begin{array}{c} d \\ m(t) \\ x(t) \\ I(t) \end{array} \end{array} = \begin{bmatrix} p1 & 0 & 1 & 0 & p2 \\ 0 & \frac{-1}{t_G} & 0 & 0 & 0 \\ 0 & \frac{1}{t_G} & \frac{-1}{t_G} & 0 & 0 \\ 0 & 0 & 0 & -k_a & 0 \\ 0 & 0 & 0 & \frac{k_a}{V_d} & -k_e \end{bmatrix} \begin{bmatrix} G(t) \\ g(t) \\ m(t) \\ x(t) \\ I(t) \end{bmatrix} + \begin{bmatrix} p_3 \\ \frac{A_G}{t_G} D_G(t) \\ 0 \\ u(t) \\ 0 \end{bmatrix}$$
 Meal Input Insulin Input

- Fit the parameters to reproduce the key glycemic statistics
 - Ranges of parameters are given in clinical literature

Population Statistics

	CSII	Model
	Dataset BG	Simulated BG
Mean BG	163	159
Max BG	365	379
Min BG	50	49
BG > 180	35%	30%
BG < 70	3%	3%
BG in [70,180]	62%	67%

Per-Subject Statistics

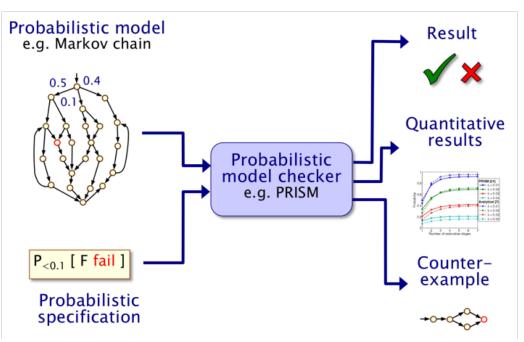
Metric	Value
Mean Difference of Per-Patient Mean BG	14 mg/dL
Mean Difference of Per-Patient $BG > 180$ Percentage	5%
Mean Difference of Per-Patient BG < 70 Percentage	1%
Mean Difference of Per-Patient BG in [70,180] Percentage	6%





Probabilistic Model Checking

- The PRISM model checker
 - The coupled system can be expressed as discrete-time Markov chains
 - Exhaustively checks all execution paths of a model against probabilistic specifications







Closed-Loop Analysis

- PRISM model checker
 - Support probabilistic transitions
 - Enables exhaustive check all execution paths of a model
- Integrate individualized physiological model and behavioral models
 - Explore how changing behavior types may impact outcomes
 - Hypoglycemia: % of CGM readings < 70 mg/dL
 - Hyperglycemia: % of CGM readings > 180 mg/dL

	ЕТС Туре	Hypoglycemia Rate (%)	Hyperglycemia Rate (%)
Actual type	E3T2C1	6.93	8.43
Change	E1T2C1	6.20	12.78
E subtype	E2T2C1	5.99	13.72
Change	E3T1C1	0.02	10.33
T subtype	E3T3C1	0.04	10.09
	E3T4C1	0.02	11.05
Change	E3T2C2	7.04	6.30
C subtype	E3T2C3	6.95	7.93
Changa	E2T1C1	0.04	16.46
Change	E2T2C1	5.99	13.72
multi-subtypes	E3T1C3	0.10	9.76
	E2T1C3	0.08	15.42

	ЕТС Туре	Hypoglycemia Rate (%)	Hyperglycemia Rate (%)
Actual type	E1T1C1	0	43.92
Change	E2T1C1	0	44.38
E subtype	E3T1C1	0	41.62
Change T subtype	E1T2C1	0	39.13
	E1T3C1	0	43.46
	E1T4C1	0	45.31
Change	E1T1C2	0	41.59
C subtype	E1T1C3	0	43.47
Change multi-subtypes	E1T2C2	0	37.22
	E3T2C1	0	35.45
	E3T1C2	0	38.01
	E3T2C2	0	32.56





Conclusion

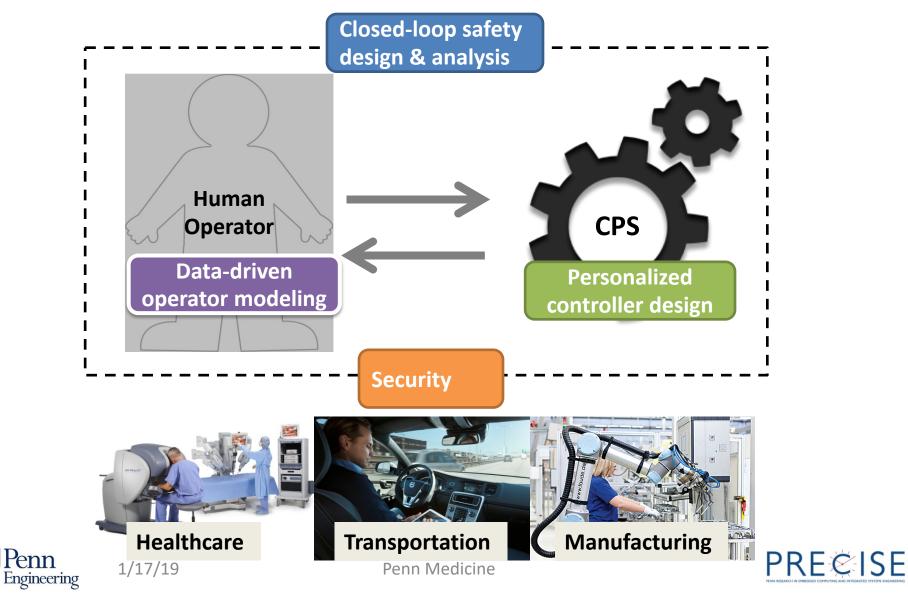
- "Eat, Trust, and Correct" behavioral modeling framework for T1D on insulin pumps
- Learn ETC behavioral clusters from clinical data
- Closed-loop analysis suggests switching behavioral types may improve glycemic control outcomes
 - More effective patient education and peer-support
- Future work
 - Testing on larger clinical datasets
 - Further development and validation of learning techniques
 - Plug in other physiological models





Challenges

Human-in/on-the-loop CPS/IoT: Modeling, design and analysis



Resources

- References
 - Challenges and Research Directions in Medical Cyber–Physical Systems, I. Lee, et al., Proceedings of the IEEE, 2012.
 - Medical Cyber-Physical Systems, I. Lee, et al., Book Chapter, June 2016.
- Publications
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- PRECISE Center
 - https://precise.seas.upenn.edu/
- Medial Device Club
 - https://rtg.cis.upenn.edu/meddevclub/
- Penn Health Tech

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– https://healthtech.upenn.edu/





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THANK YOU!



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