

# DISH-TREND: INTERVENTION MODELING SIMULATOR THAT ACCOUNTS FOR TREND INFLUENCES

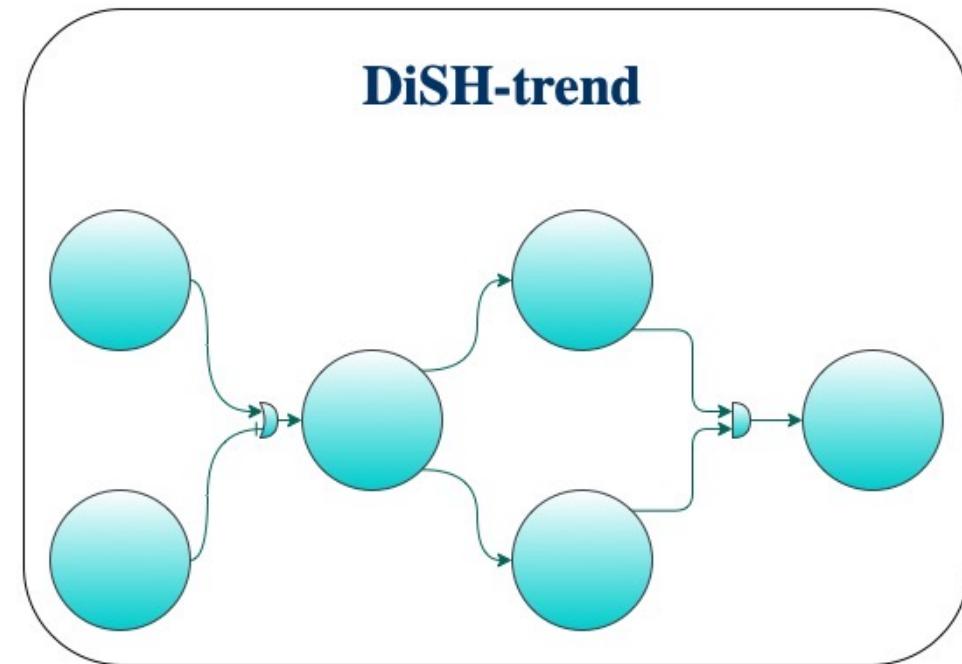
WINTER SIMULATION CONFERENCE 2021

STEFAN ANDJELKOVIC & NATASA MISKOV-ZIVANOV



# INTRODUCTION

- Dynamic network models – crucial in understanding complex system dynamics
- Nodes = components, edges = interactions
- Applications: biology, psychology, sociology, economics, politics, and engineering
- Interventions = constraints on values, or structural modifications



# SIMILAR MODELING APPROACHES

- Boolean networks (Albert et al 2008)
- Discrete networks (Sayed et al 2017)
- Bayesian networks (Needham 2007, Wilkinson et al 2007)
- ODE-based models (Materi and Wishart 2007)
- Rule-based models (Faeder et al 2005)
- Petri nets (Chaouiya et al 2007)

# DISH (DISCRETE STOCHASTIC HETEROGENEOUS) SIMULATOR

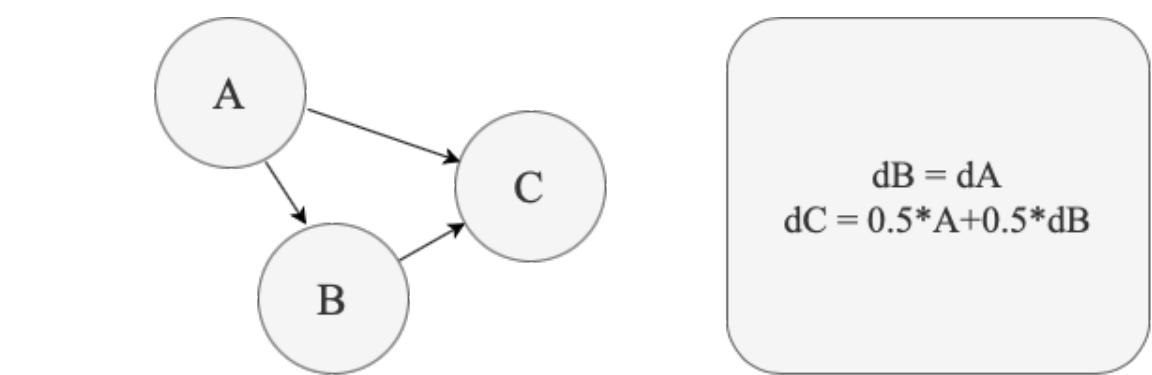
- Discrete levels (extension from Boolean networks)
- Update functions (Boolean)
- BioRECIPES model representation: element-focused
- Simulation schemes: simultaneous vs sequential
- Value toggling (enforced value constraints)

# FROM DISH TO DISH-TREND

- Algebraic update function: weighted sums of regulation functions
  - $B(\mathbf{A}, \mathbf{I}) = \sum_{i=1}^k \left( w_i^{(p)} \prod_{q \in \mathbb{I}_{p,i}} A_{i,q} \right) - \sum_{j=1}^l \left( w_j^{(n)} \prod_{r \in \mathbb{I}_{n,j}} I_{j,r} \right)$  (WEIGHTED SUM-OF-PRODUCTS)
  - $F(\mathbf{A}, \mathbf{I}) = \frac{\lceil L \cdot |B(\mathbf{A}, \mathbf{I})| \rceil}{L} \cdot \text{sign}(B(\mathbf{A}, \mathbf{I}))$
- Adding trend-based regulation
  - $B_{trend}(\mathbf{A}, \mathbf{I}) = \sum_{i=1}^k \left( w_i^{(p)} \prod_{q \in \mathbb{I}_{p,i}} \Delta A_{i,q} \right) - \sum_{j=1}^l \left( w_j^{(n)} \prod_{r \in \mathbb{I}_{n,j}} \Delta I_{j,r} \right)$
- Hybrid regulation – the sum of level-based and trend-based regulation
- Beyond linear models – any polynomial of  $A_{i,q}$ ,  $\Delta A_{i,q}$ ,  $I_{j,r}$ , and  $\Delta I_{j,r}$

# SMALL EXAMPLE

- 3-element network
- A – controlled input
- $t=1$ : B remains unchanged
- $t=4$ : B increased by  $A(t=3) - A(t=0)$



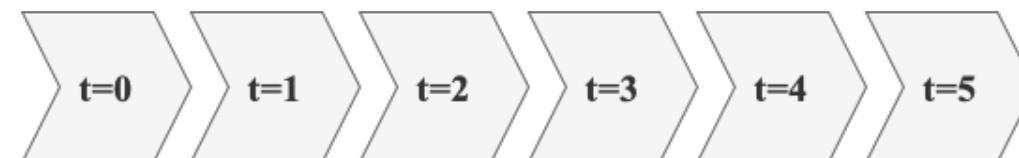
Toggle sequence of A:  
 $v=0.2, (v=0.6, t=2), (v=0.4, t=3)$

Update sequence:  
B, C, C, B, C

A:	0.2	0.2	0.6	0.4	0.4	0.4
----	-----	-----	-----	-----	-----	-----

B:	0.1	0.1	0.1	0.1	0.3	0.3
----	-----	-----	-----	-----	-----	-----

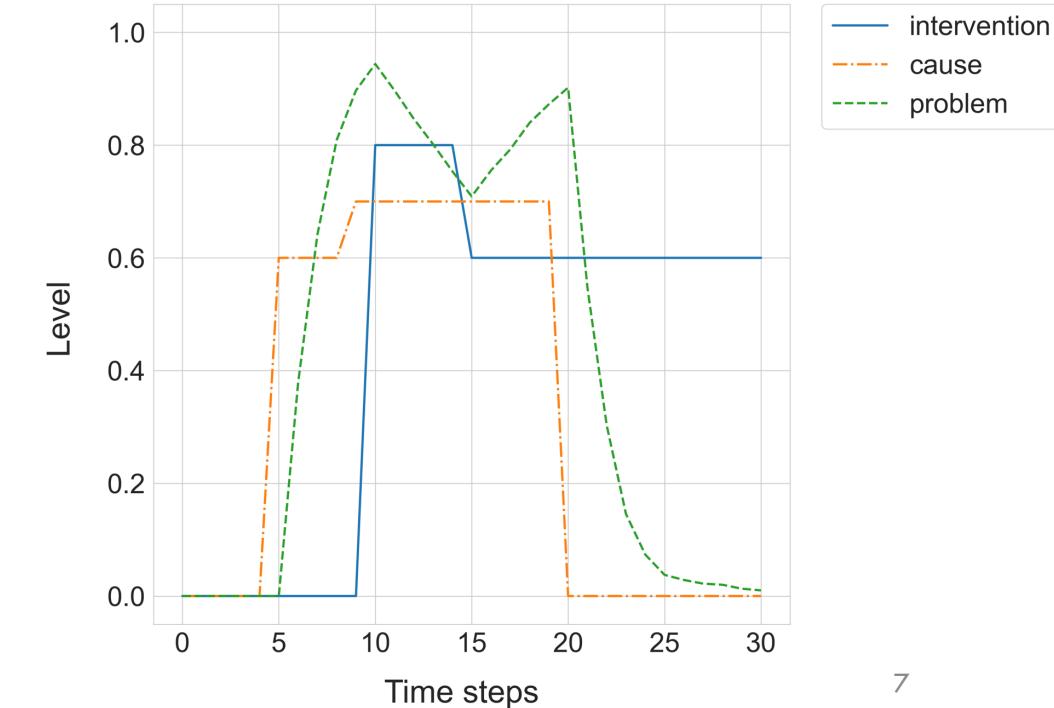
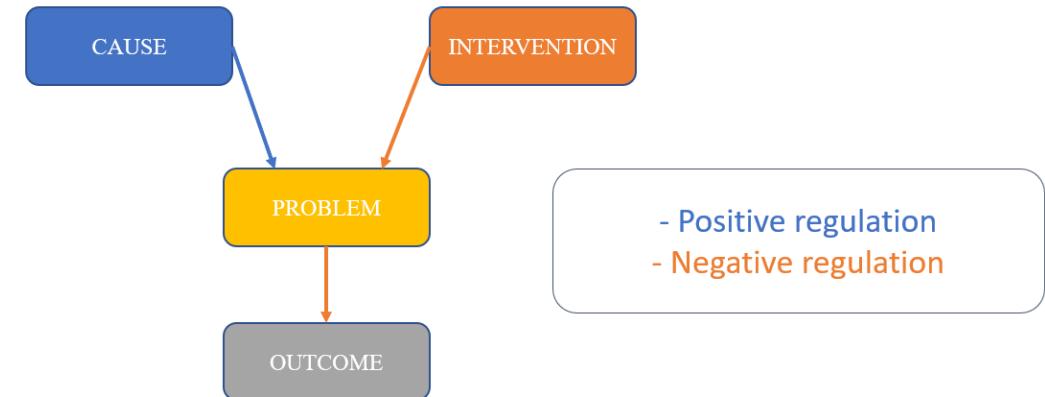
C:	0	0	0.1	0.4	0.4	0.7
----	---	---	-----	-----	-----	-----



# INTERVENTION TOY MODEL

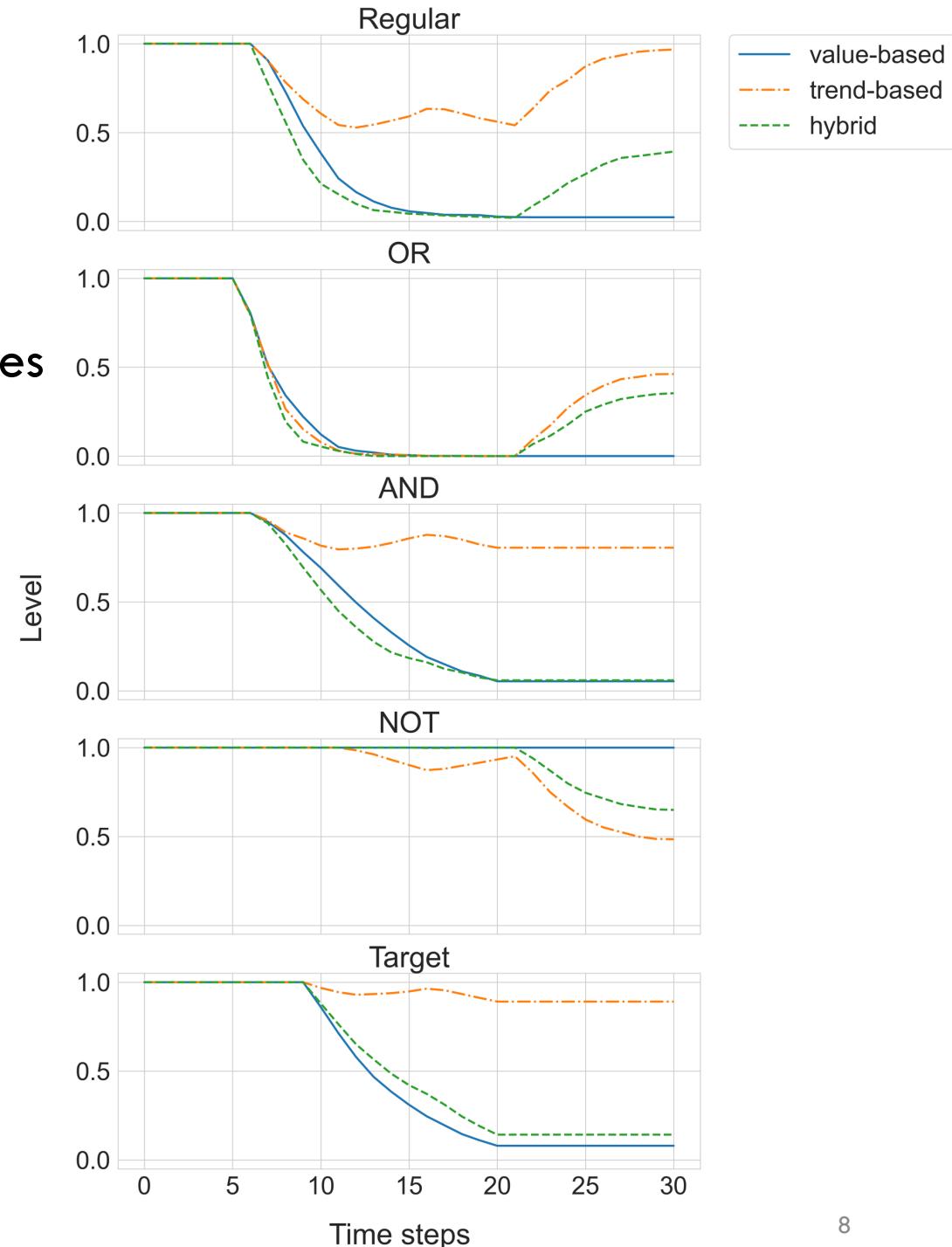
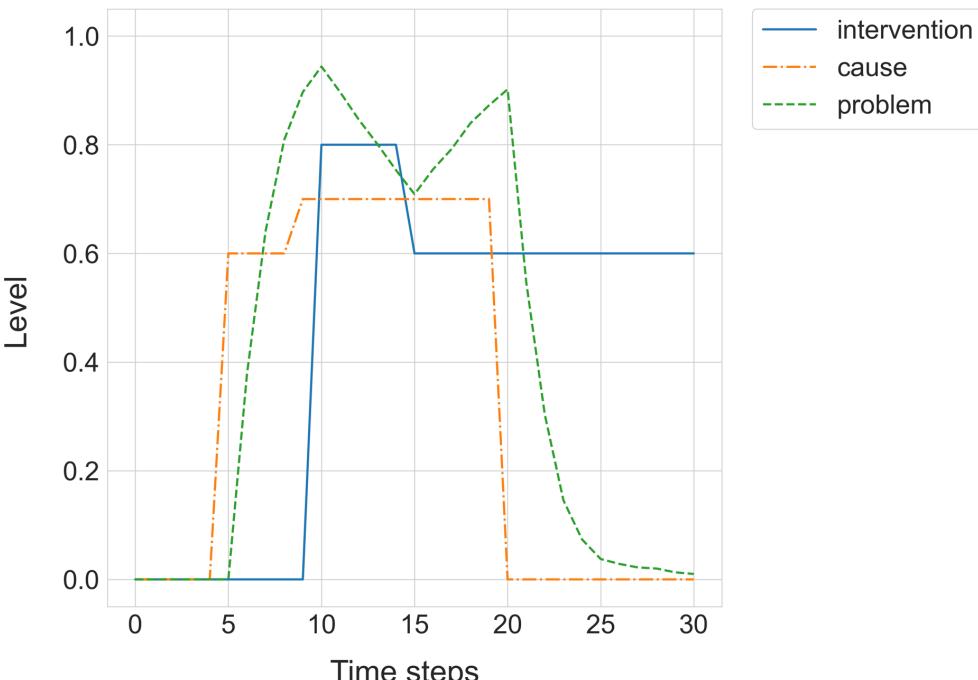
- 4-element network
- Linear model
- 5 tested scenarios:

Scenario	Negative regulation
Regular	$d(\text{OUTCOME}) = -\text{PROBLEM}$
OR	$d(\text{OUTCOME}) = -(\text{PROBLEM} + \text{CAUSE})$
AND	$d(\text{OUTCOME}) = -(\text{PROBLEM} * \text{CAUSE})$
NOT	$d(\text{OUTCOME}) = \text{PROBLEM}$
Target	$d(\text{OUTCOME}) = -(\text{PROBLEM} * \text{CAUSE})$ only when $\text{CAUSE}=0.7$



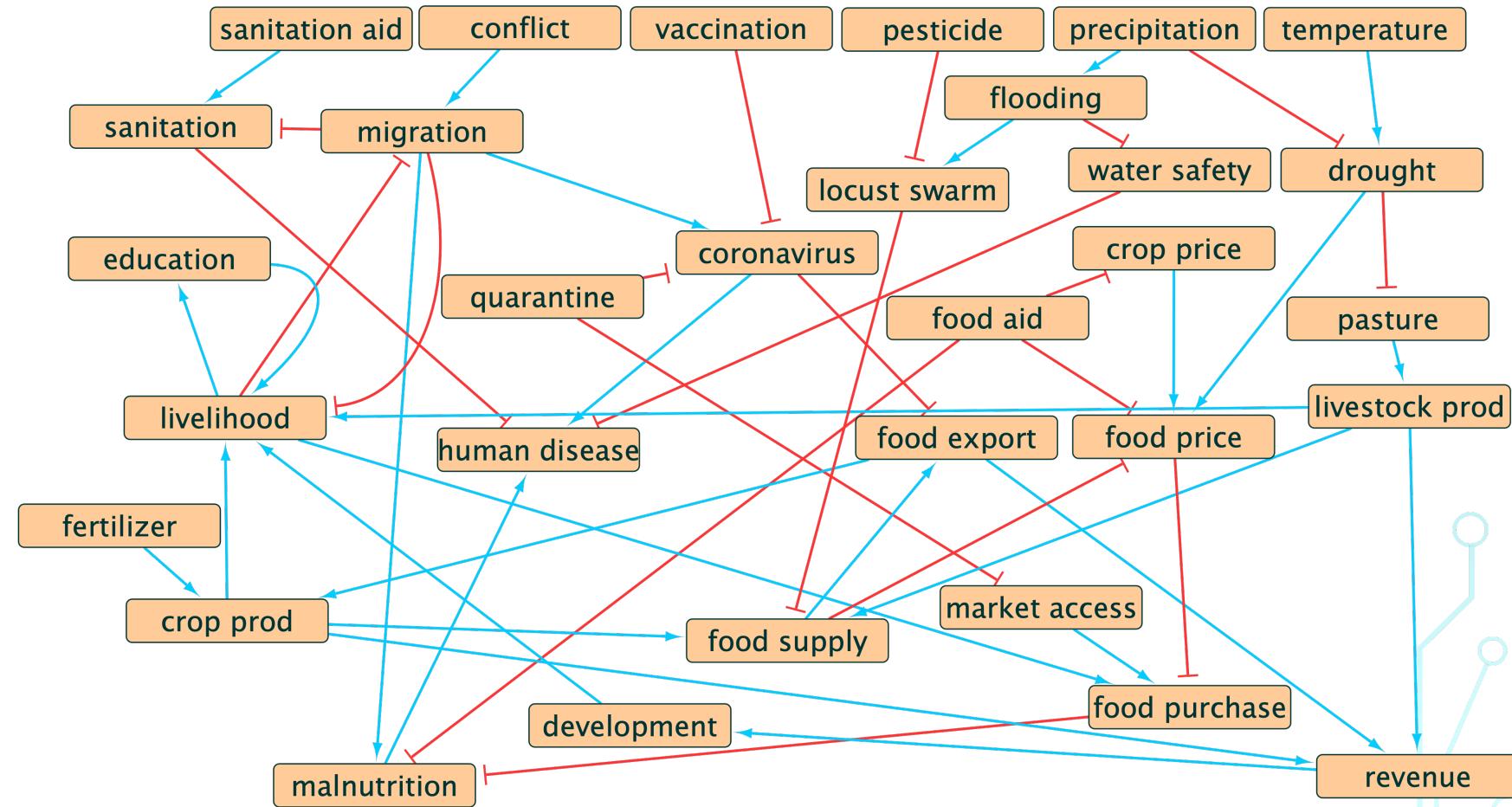
# RESULTS

- Toy model, 5 scenarios, 3 regulation types
- Execution time: 0.37-0.54 s



# LARGER MODEL – FOOD INSECURITY IN ETHIOPIA

- 31 nodes, 49 edges
- 10 inputs:
  - Temperature
  - Precipitation
  - Pesticide
  - Medical treatment
  - Fertilizer aid
  - Sanitation aid
  - Food Aid
  - Conflict
  - Quarantine
  - Vaccination

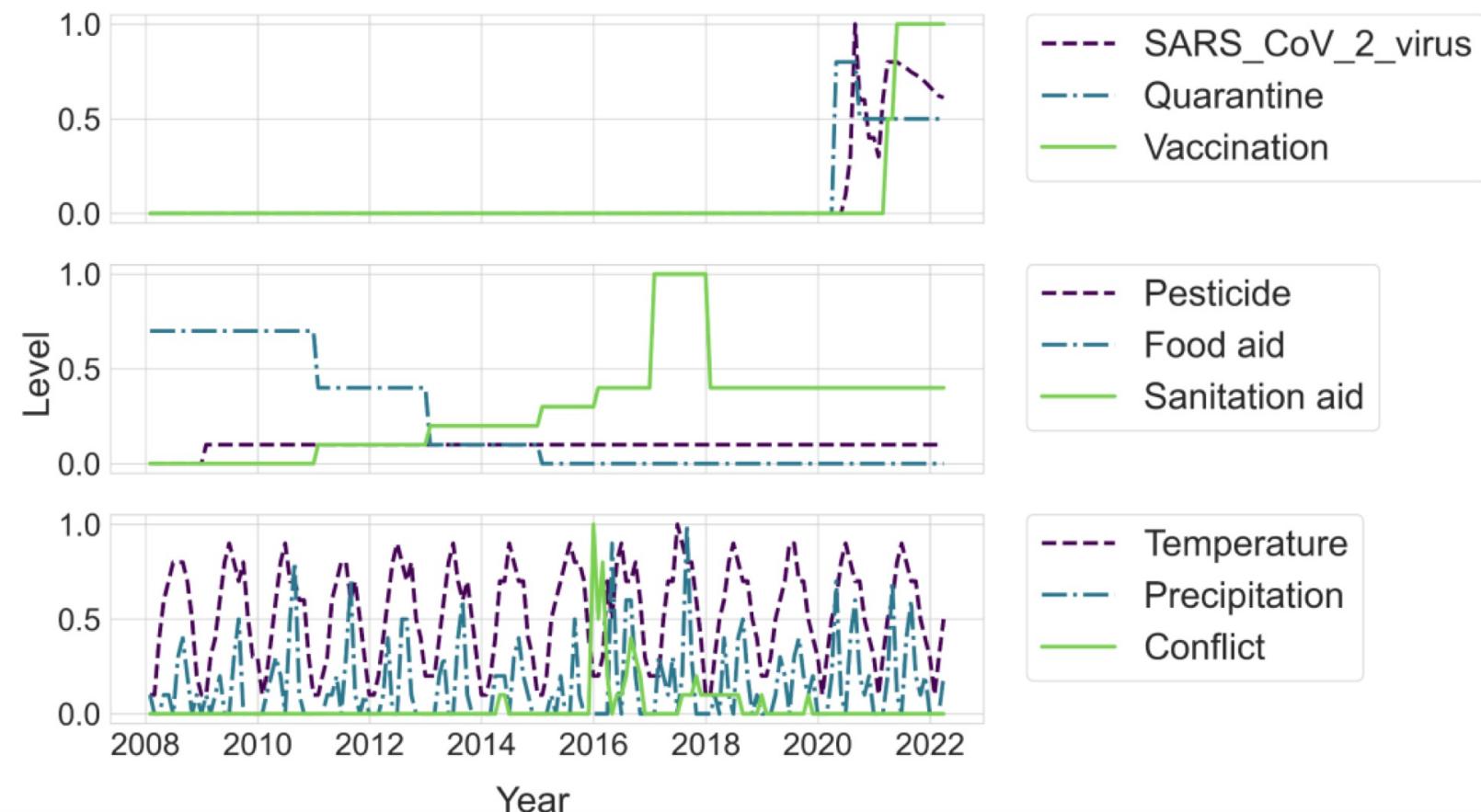


# LARGE MODEL – DATA SOURCES

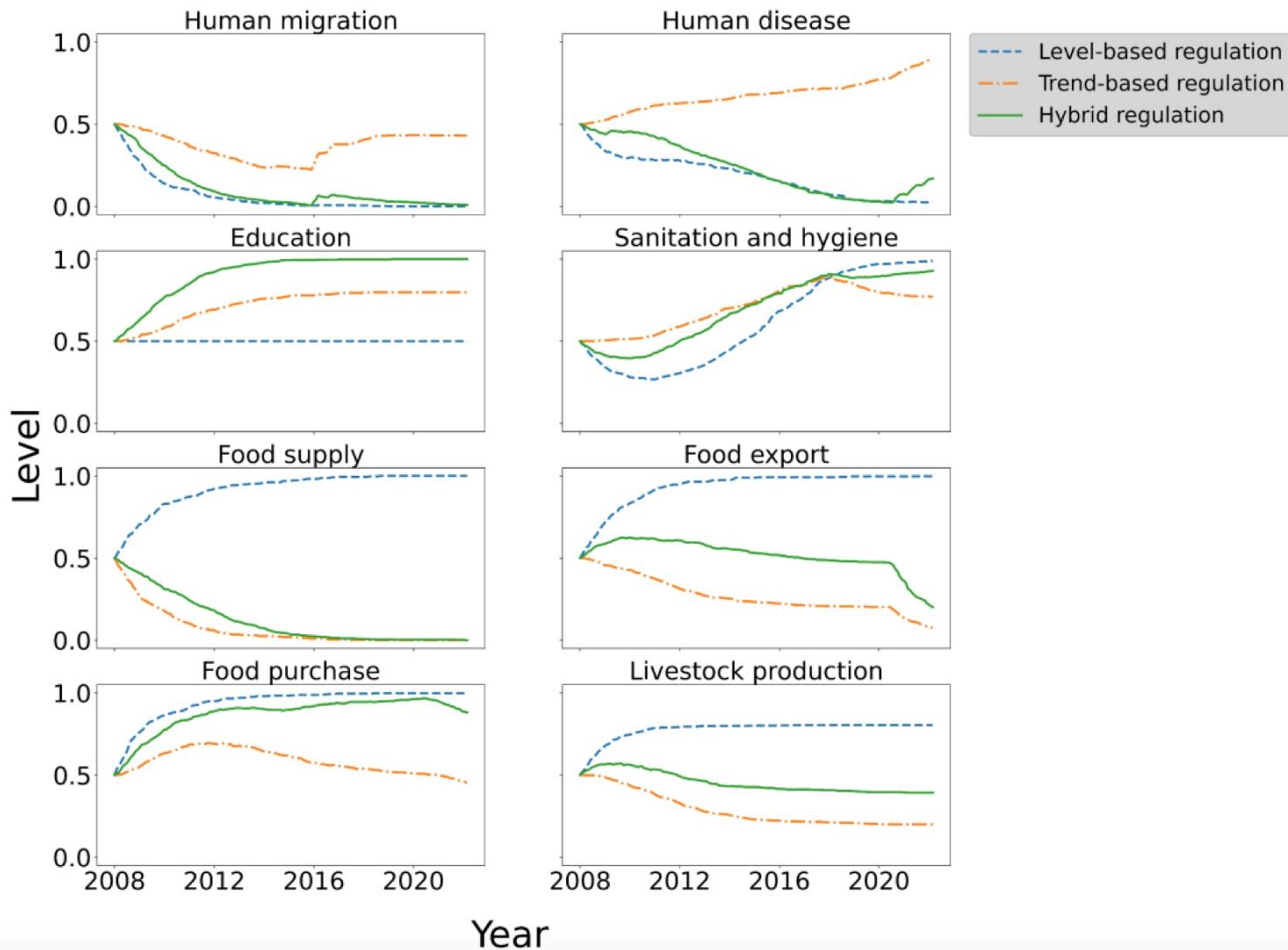
Variable	Source	Reference	Resolution (spatial and temporal)
Conflict	ACLED	Raleigh et al 2010	Ethiopia/daily
Food aid	FAOSTAT	FAOSTAT	Ethiopia/yearly
Pesticide	FAOSTAT	FAOSTAT	Ethiopia/yearly
Precipitation	CHIRPS	Funk et al 2005	Oromia/daily
Quarantine	EPHI	EPHI	Ethiopia/monthly
Sanitation and hygiene aid	USAID	USAID Database	Ethiopia/yearly
SARS-CoV-2 virus	OWID	OWID; Roser et al 2020	Ethiopia/daily
Temperature	ERA5	ERA5; Hersbach et al 2020	Oromia/daily
Vaccination	WHO	WHO	Ethiopia/daily

# RESULTS – LARGE MODEL

- Execution time:
  - Level-based: 68.7 s
  - Trend-based: 71.2 s
  - Hybrid: 74.7 s



# RESULTS – LARGE MODEL



# CONCLUSIONS

- Trend-based regulation resolves some of the level-based counter-intuitive behaviors
- Hybrid regulation gives more modeling flexibility
- DiSH-trend simulator can be used to model multi-domain complex systems
- Major advantages:
  - Highly tunable
  - Robust to missing data
  - Easy to automate and integrate with other software

# ACKNOWLEDGEMENTS

## MeLoDy lab

Mechanisms and Logic of  
Dynamics



**Natasa**

**Miskov-Zivanov, PhD**



**Kara Bocan,**  
PhD



**Cheryl**  
Telmer, PhD



**Yasmine**  
**Ahmed**



**Casey**  
**Hansen**



**Niteesh**  
Sundaram



**Evan**  
Becker



**Khaled**  
Sayed, PhD



**Adam**  
Butchy



**Emilee**  
**Holtzapple**



**Gaoxiang**  
**Zhou**



**Handa**  
Ding

## Collaborators

- George Wittenberg, MD, PhD, Neurology, UPitt
- Peter Spirtes, PhD, Philosophy, CMU
- Brent Cochran, PhD, School of Medicine, Tufts
- Sandra Cascio, PhD, Immunology, UPitt
- Michael Lotze, MD, UPCI, UPitt
- Christof Kaltenmeier, MD, UPCI, UPitt
- Ed Hovy, PhD, LTI, CMU
- Robert Stephens, PhD, NCI, NIH

<https://www.nmzlab.pitt.edu>



# ACKNOWLEDGEMENTS

- MeLoDy Lab (N. Miskov-Zivanov lab)
- Kara Bocan and Khaled Sayed  
(ex lab members)
- Collaborators: Brad Goodman, Robyn Kozierok, Zoe Henschield, Lynette Hirschman, Pam Bhattacharya, and Pascale Proulx
- DARPA grant W911NF-18-1-0017
- WSC conference