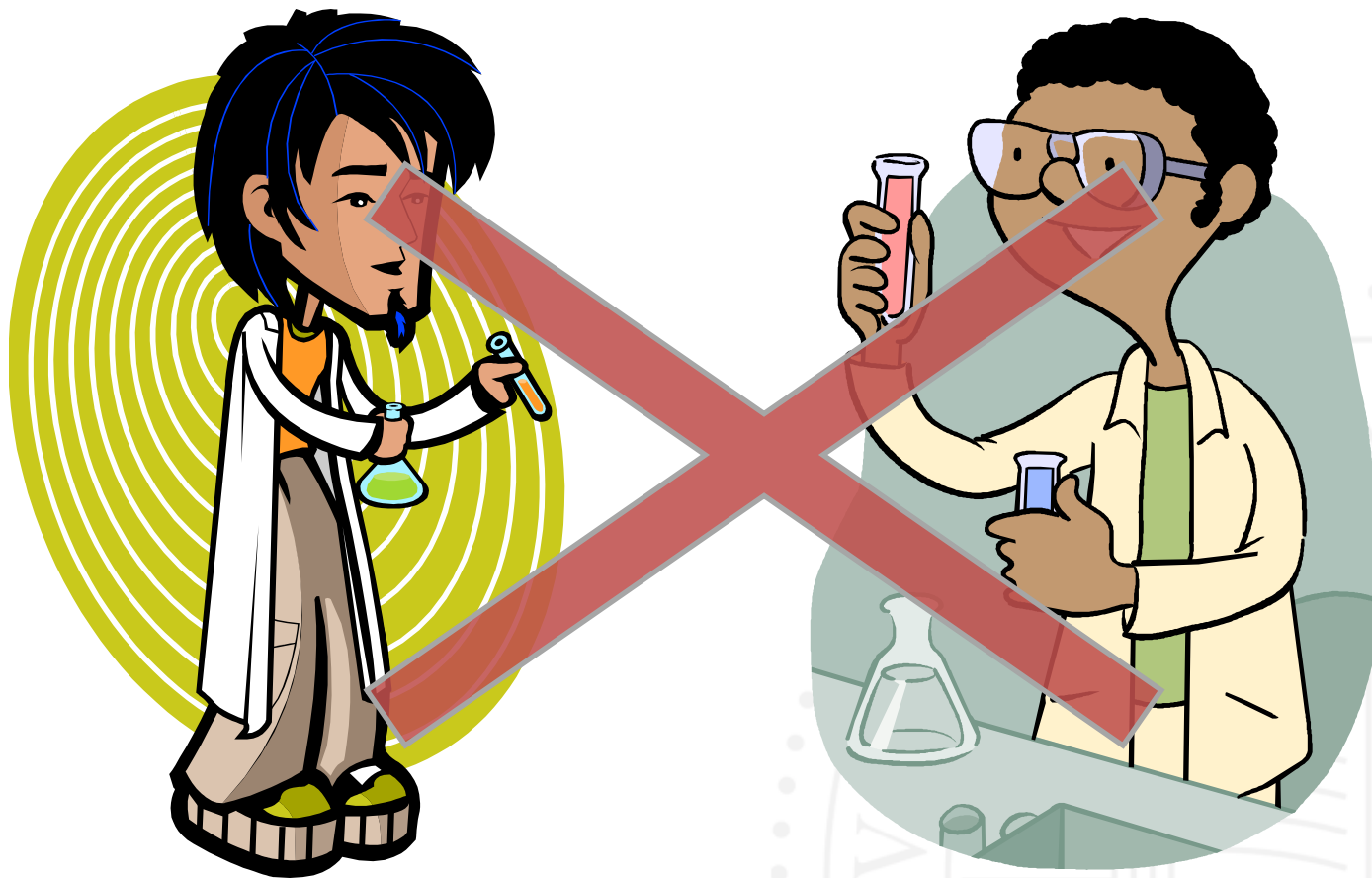


Exploring Speed and Energy Tradeoffs in Droplet Transport for Digital Microfluidic Biochips

Johnathan Fiske, ***Dan Grissom**, Philip Brisk
University of California, Riverside

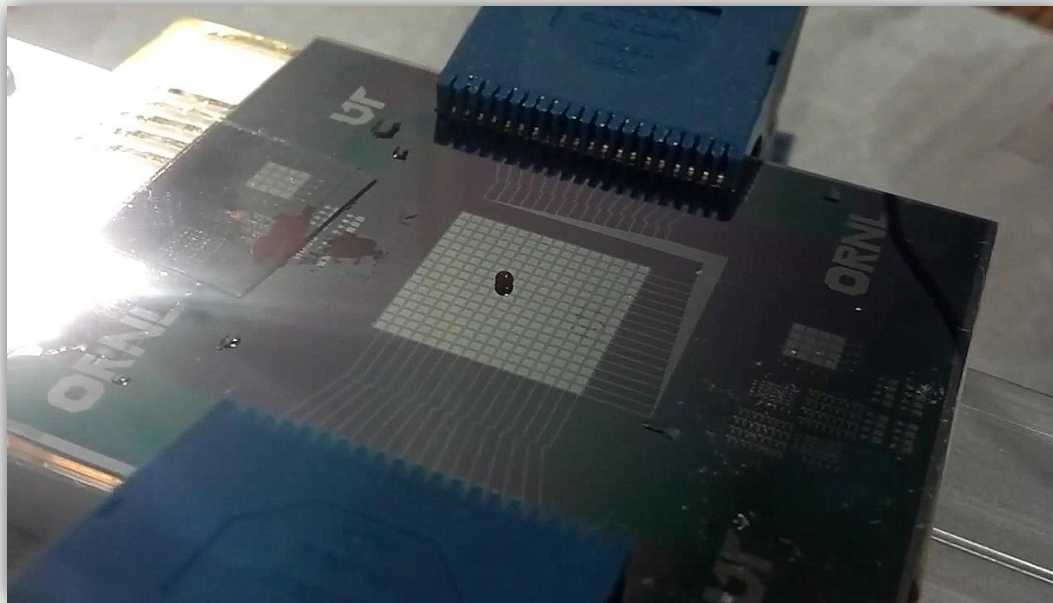
The Bottom Line



Microfluidics will replace traditional bench-top chemistry

Microfluidics

“Digital”



Discrete Droplet Based

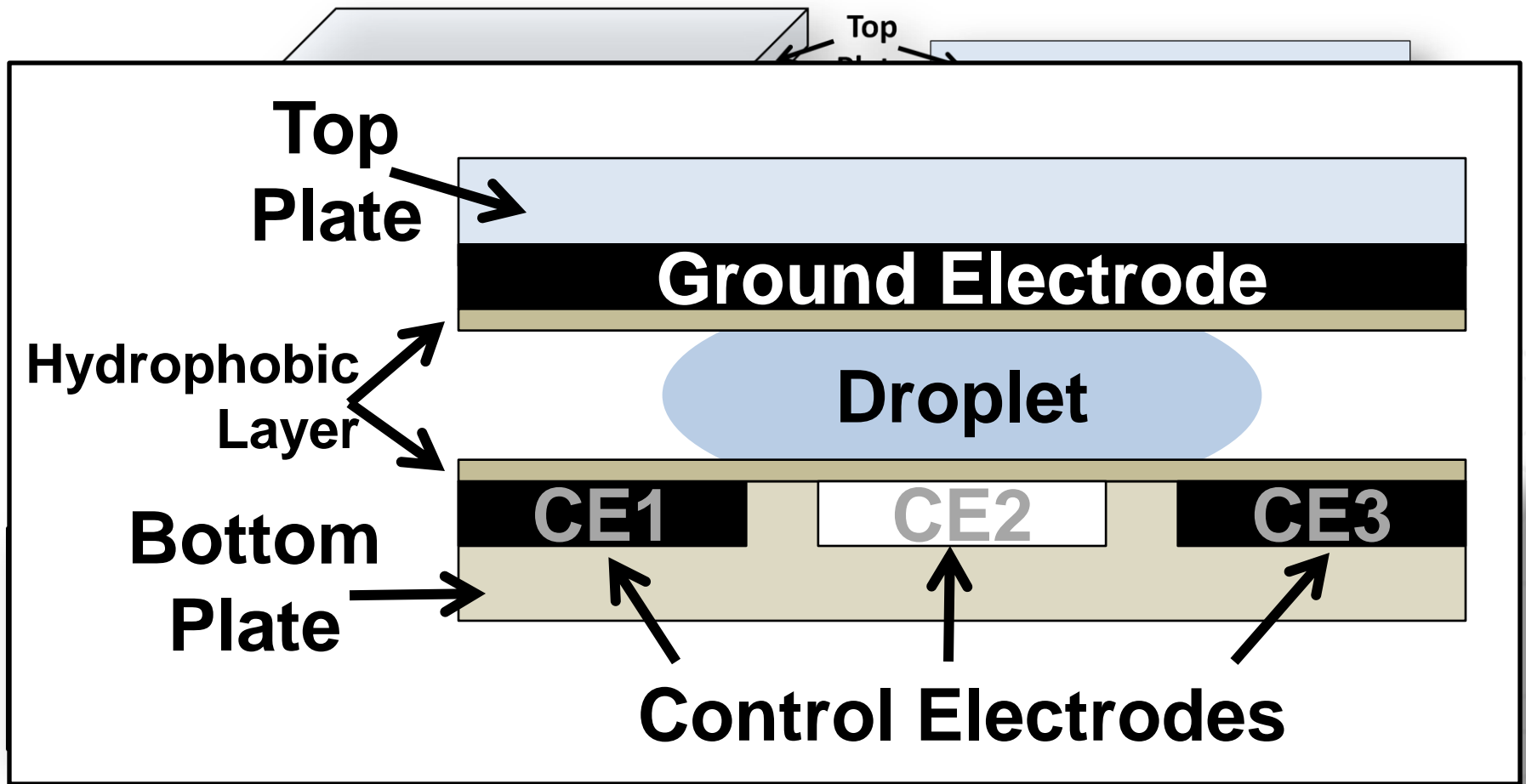
Miniaturization + Automation
of Biochemistry

Applications

- › Biochemical reactions and immunoassays
 - › Clinical pathology
- › Drug discovery and testing
 - › Rapid assay prototyping
- › Biochemical terror and hazard detection
- › DNA extraction & sequencing



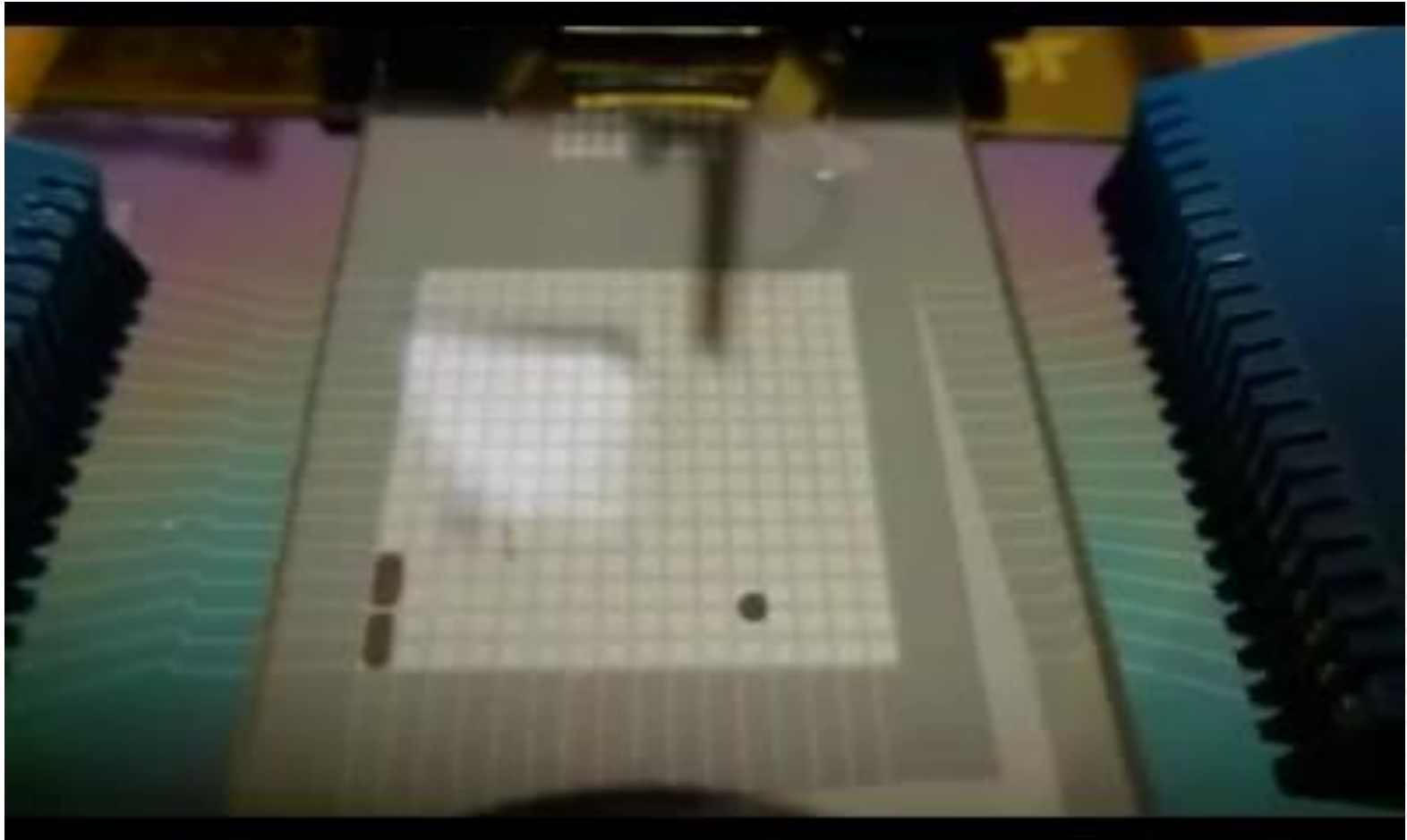
Digital Microfluidic Biochips (DMFB) 101



Basic Microfluidic Operations

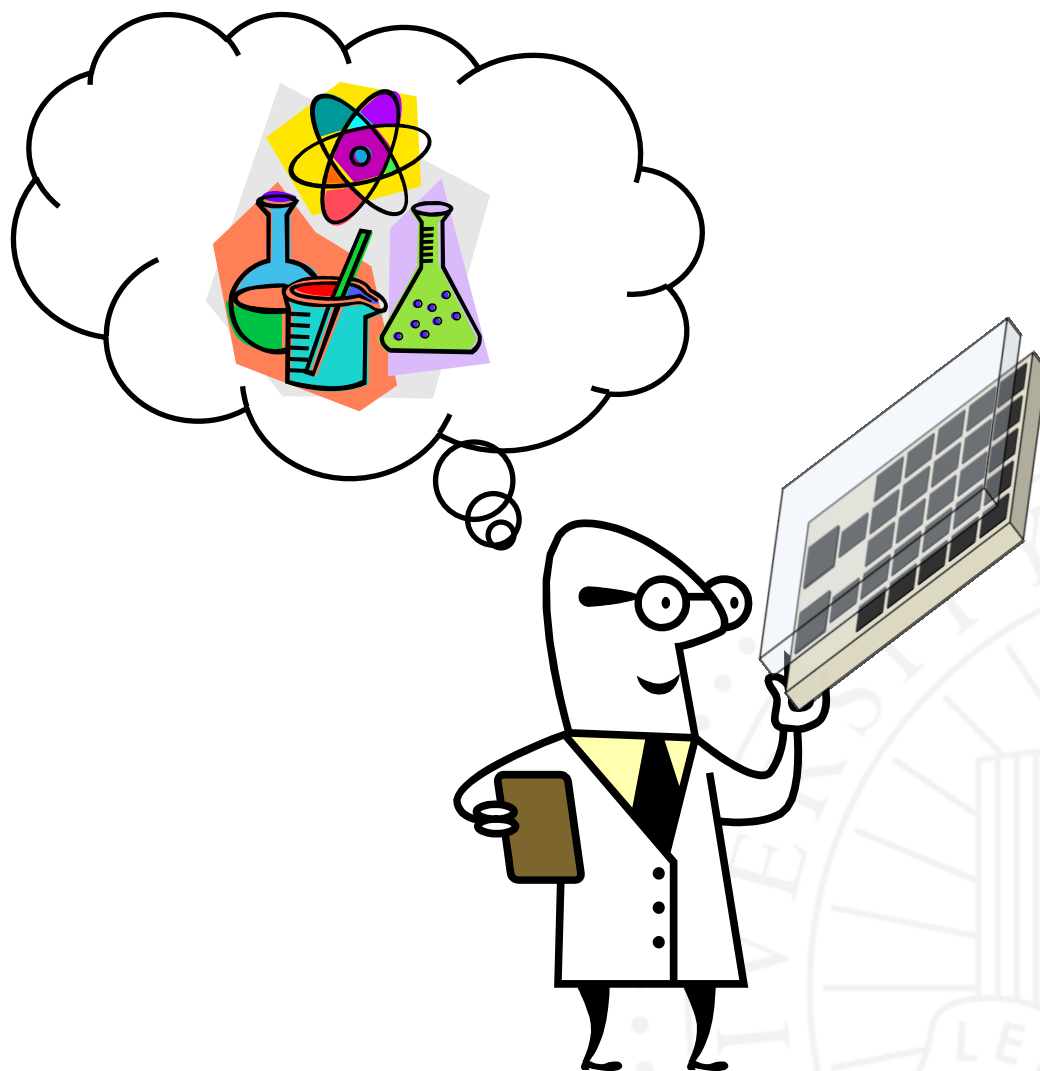
<http://microfluidics.ee.duke.edu/>

Digital Microfluidic Biochips (DMFB) 101



**Droplet Actuation on a Prototype DMFB
at the University of Tennessee**

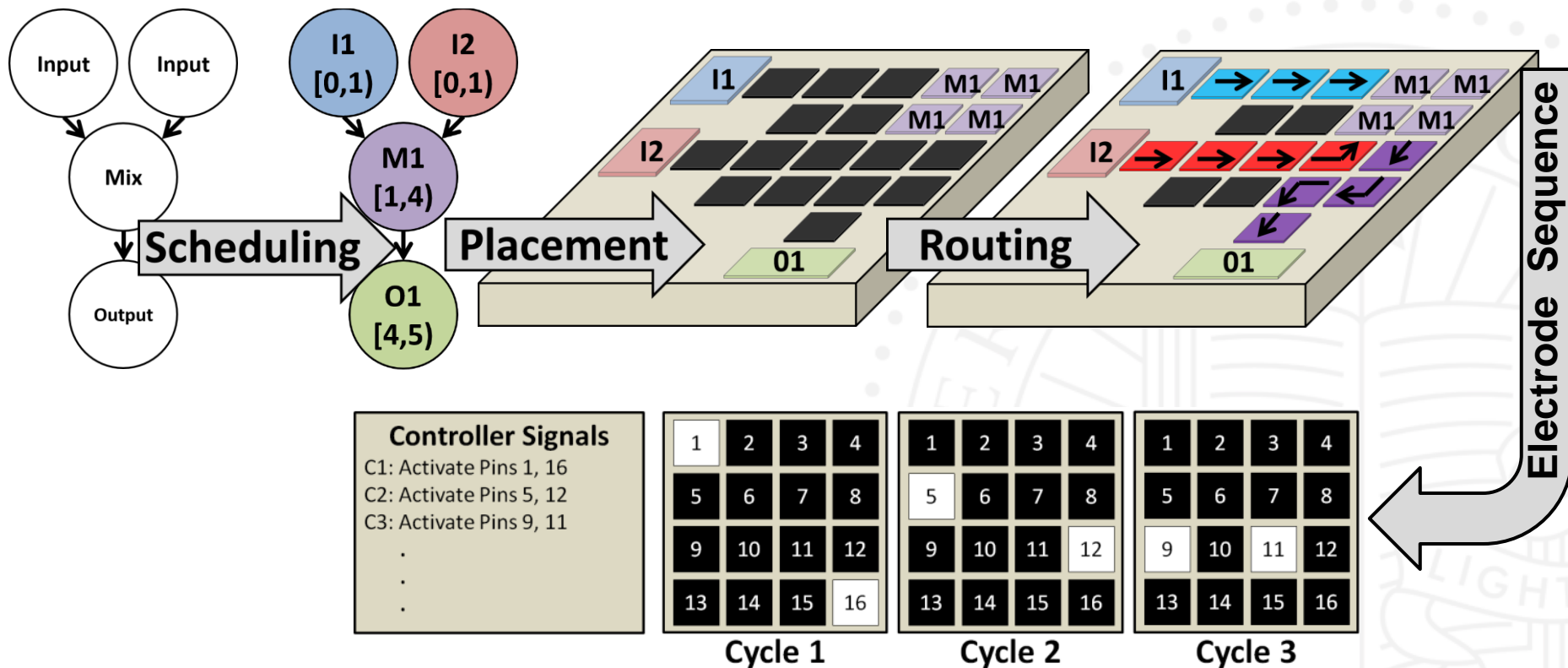
DMFB Mapping



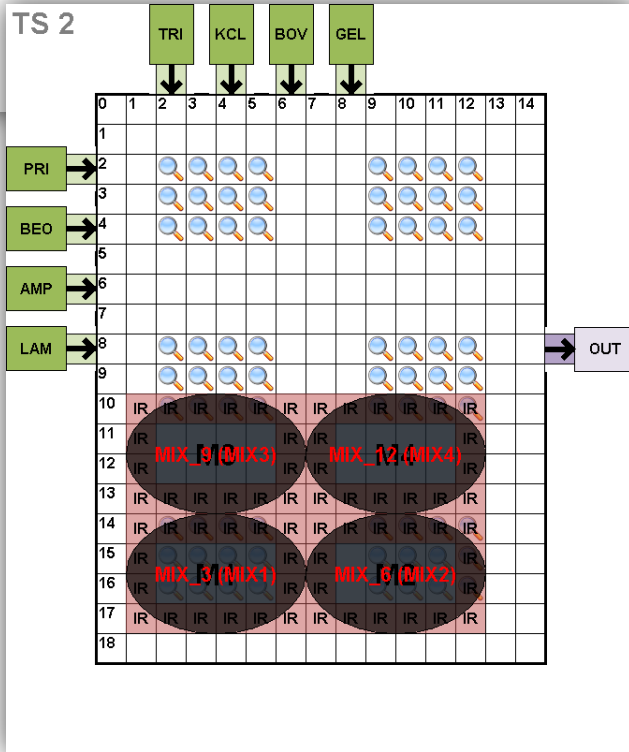
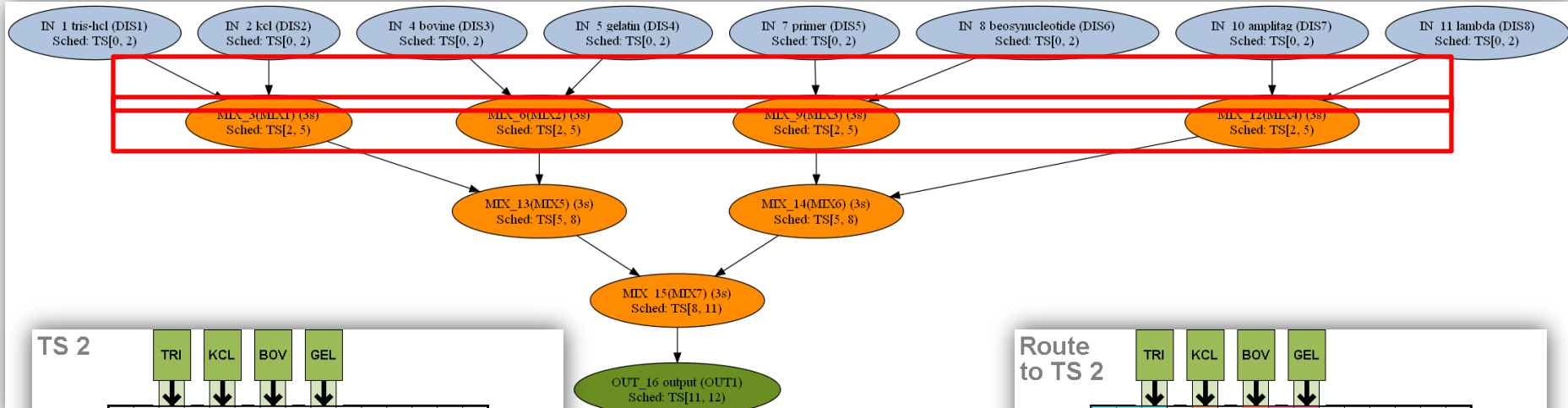
How do I make a reaction run on a DMFB?

CAD Synthesis Flow

- Synthesis: The process of **mapping** an application to hardware
 - Similar to how applications are mapped to ICs

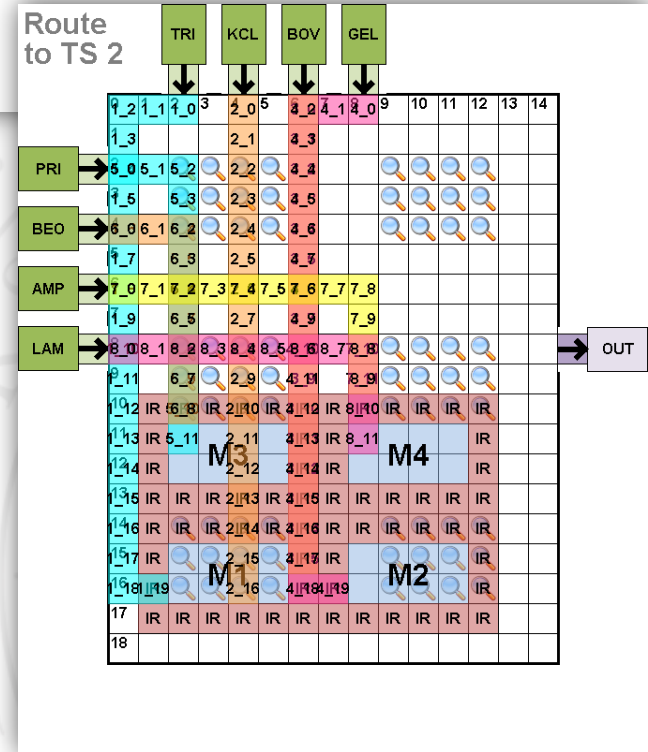


Synthesis Example



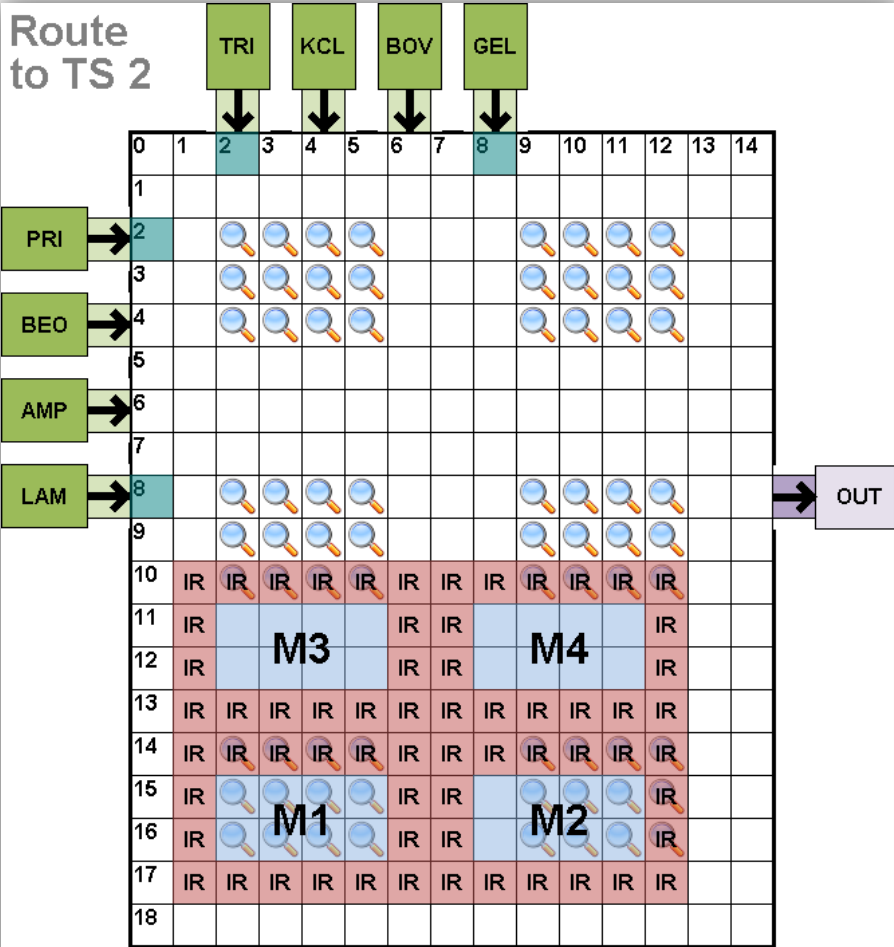
2.) Place

1.) Schedule

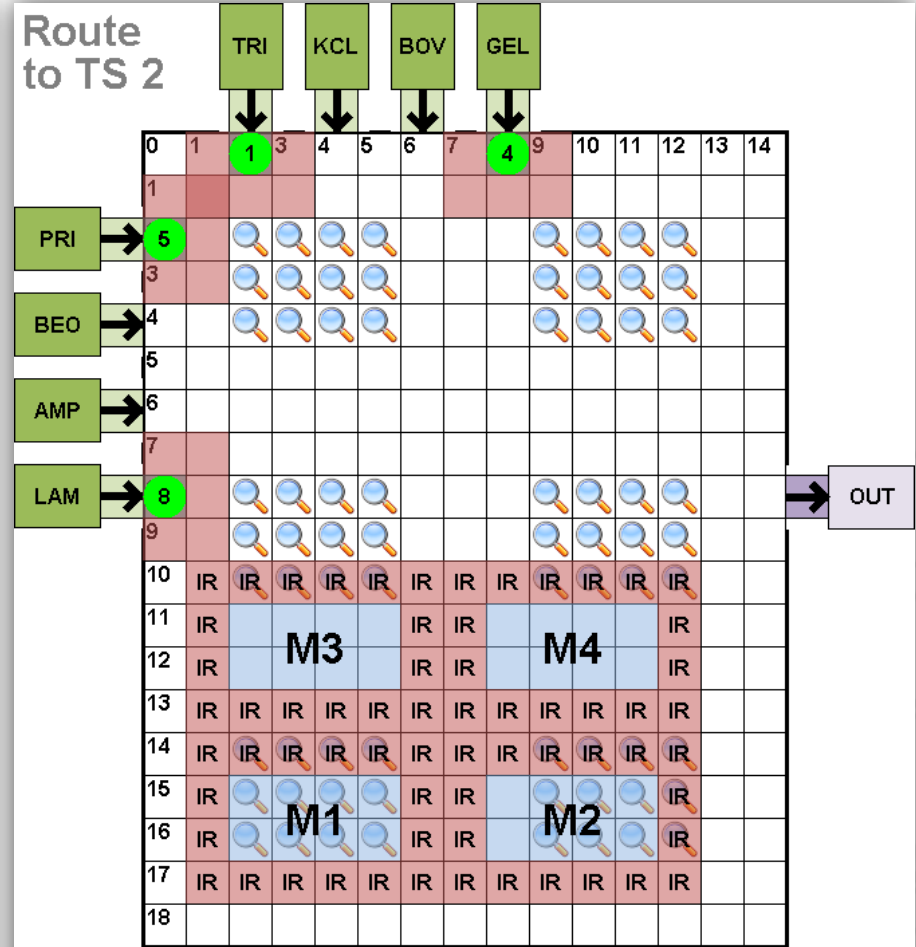


3.) Route

Compaction Example

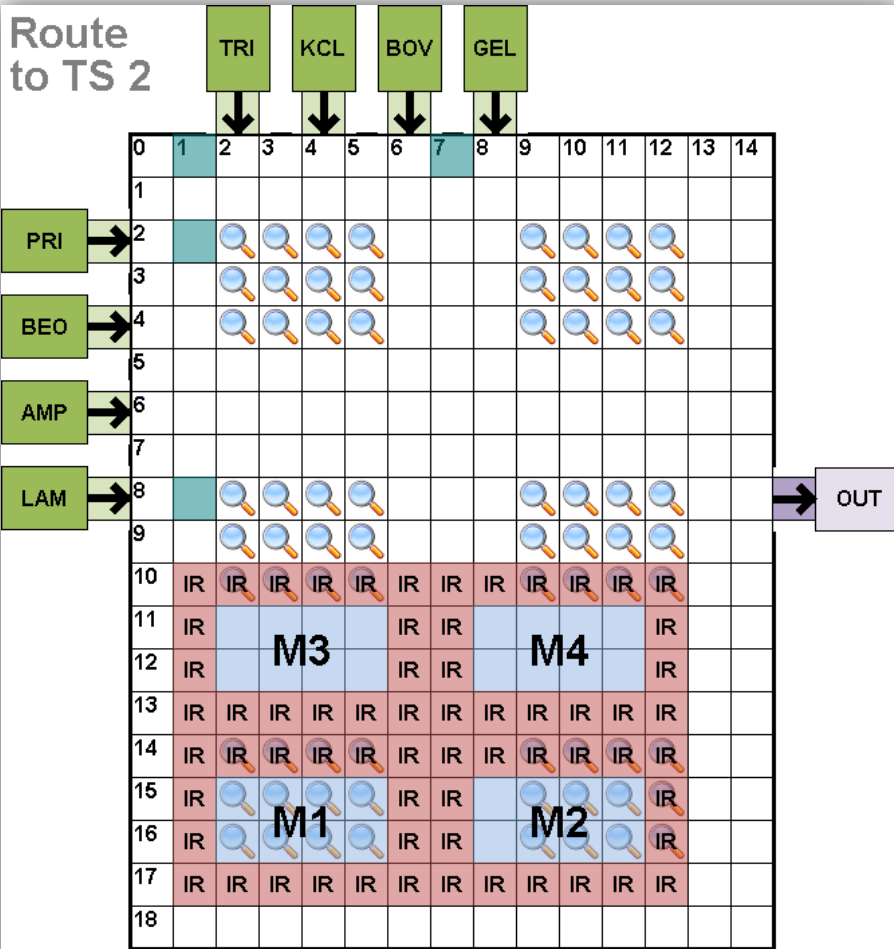


Electrode Activations

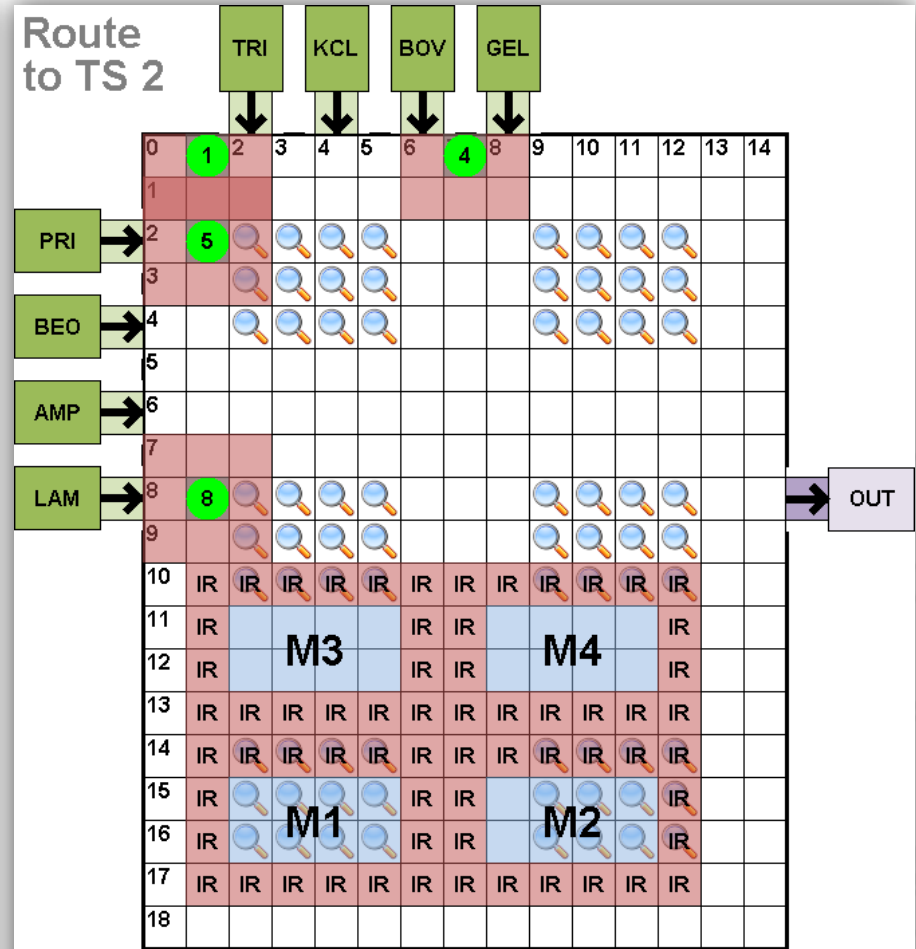


Corresponding Droplet Motion

Compaction Example

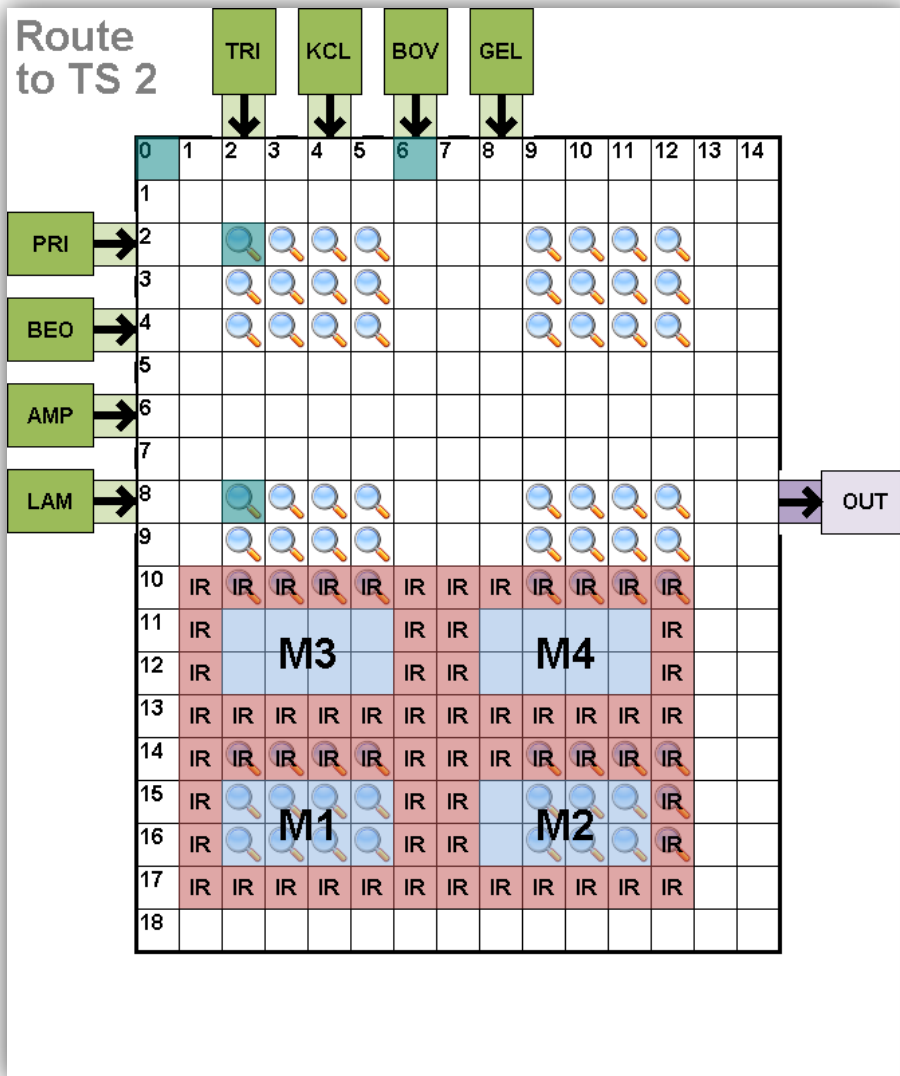


Electrode Activations

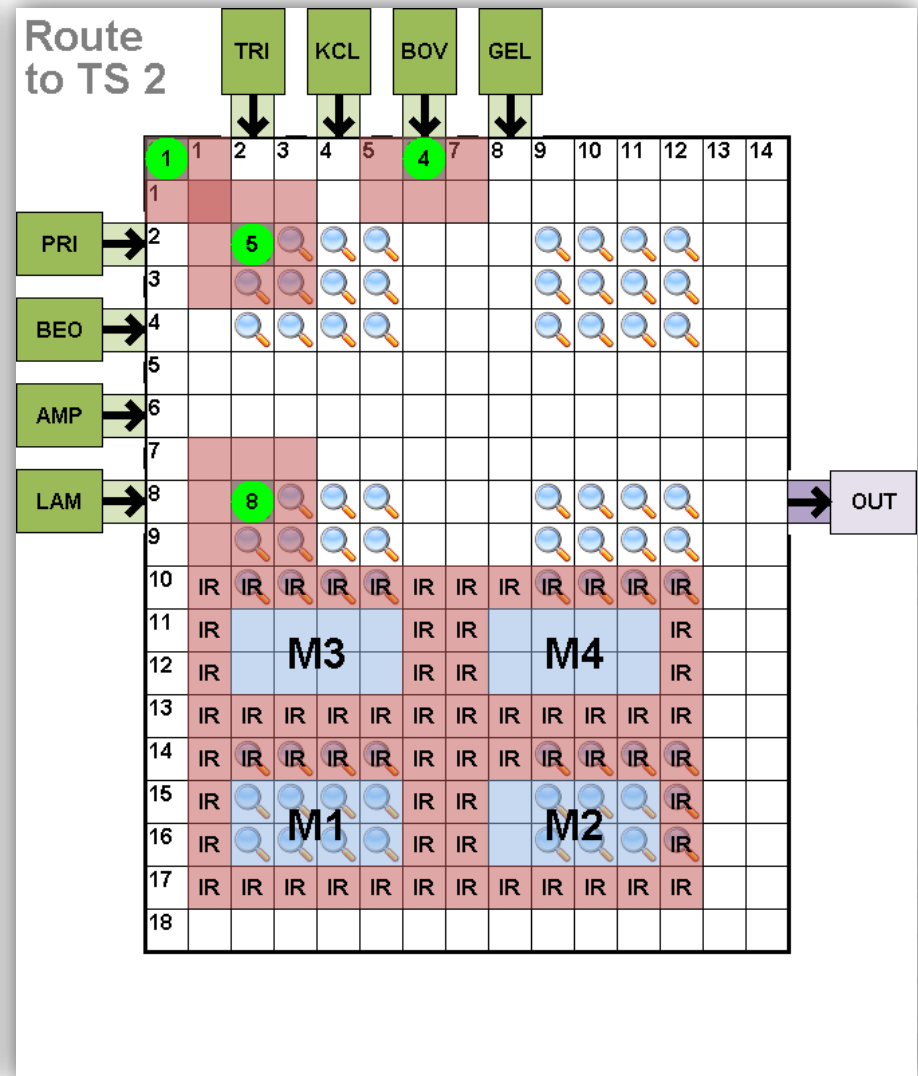


Corresponding Droplet Motion

Compaction Example

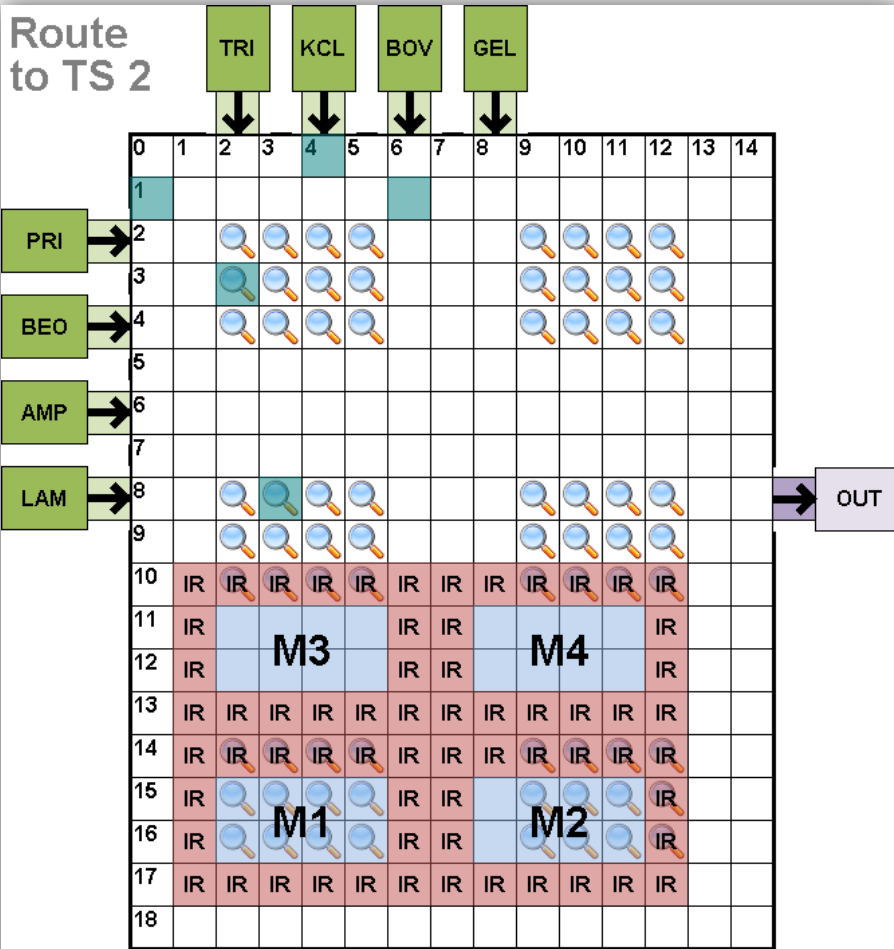


Electrode Activations

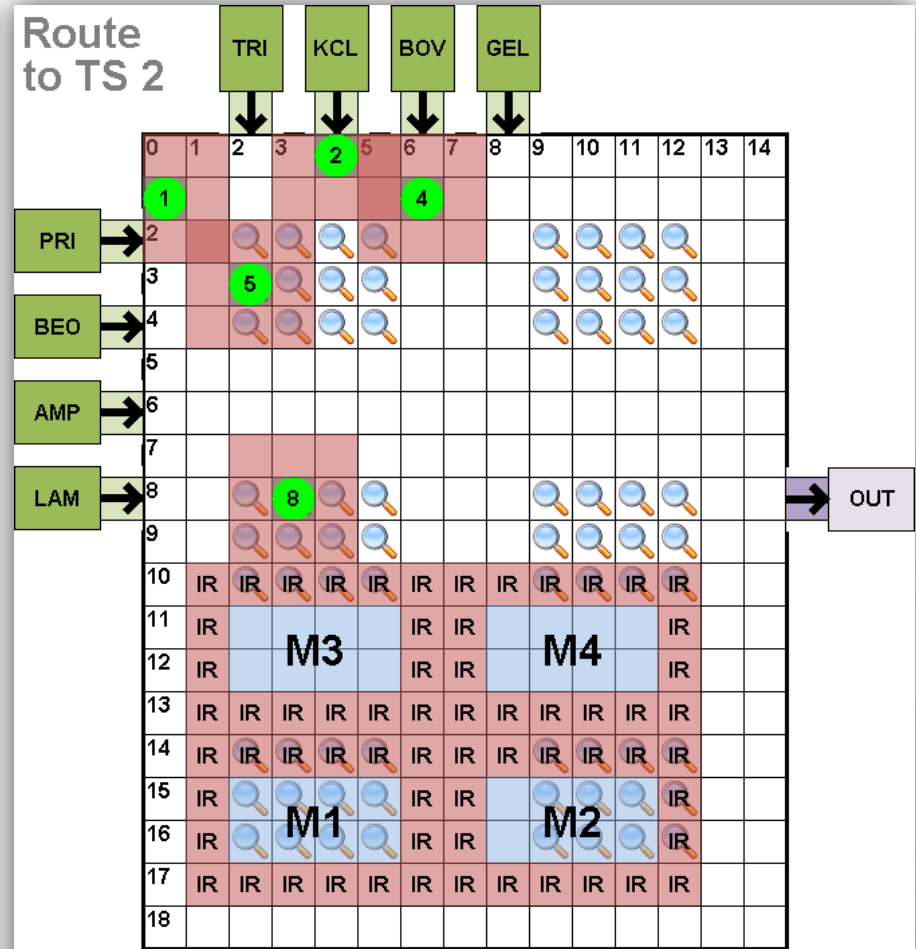


Corresponding Droplet Motion

Compaction Example

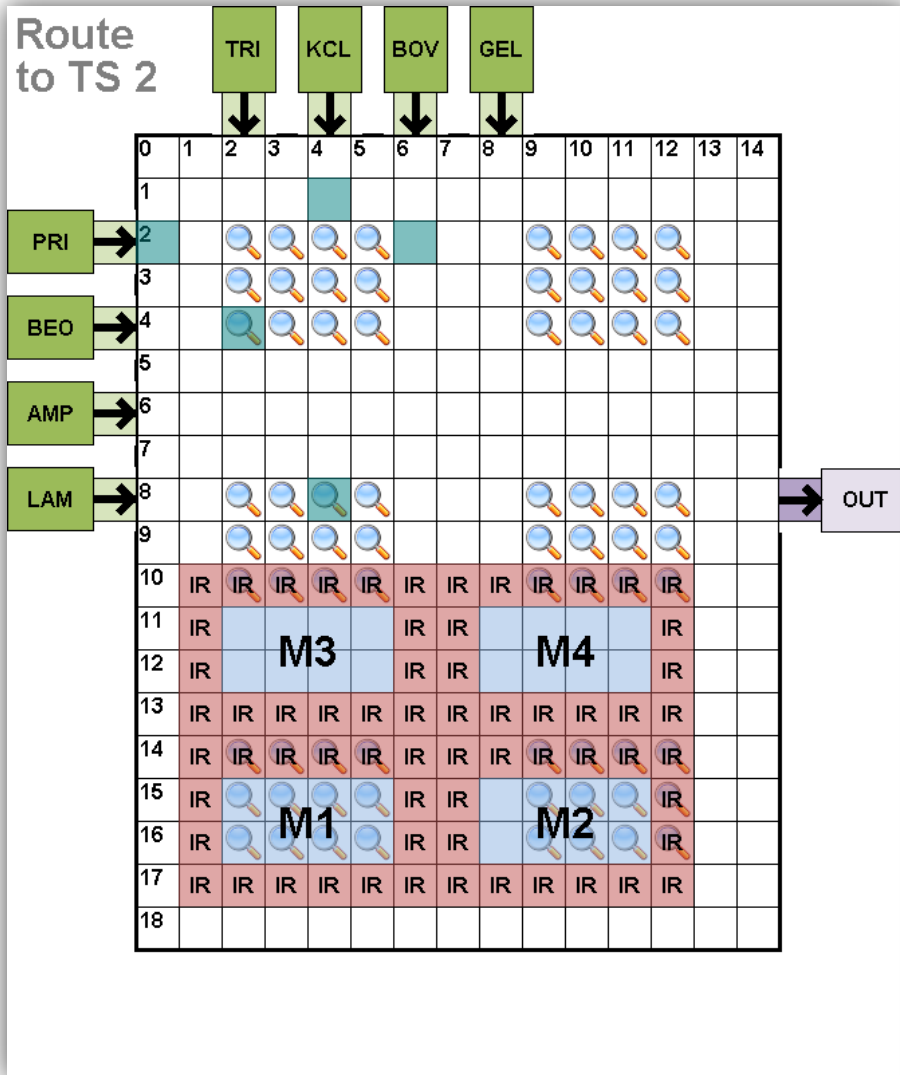


Electrode Activations

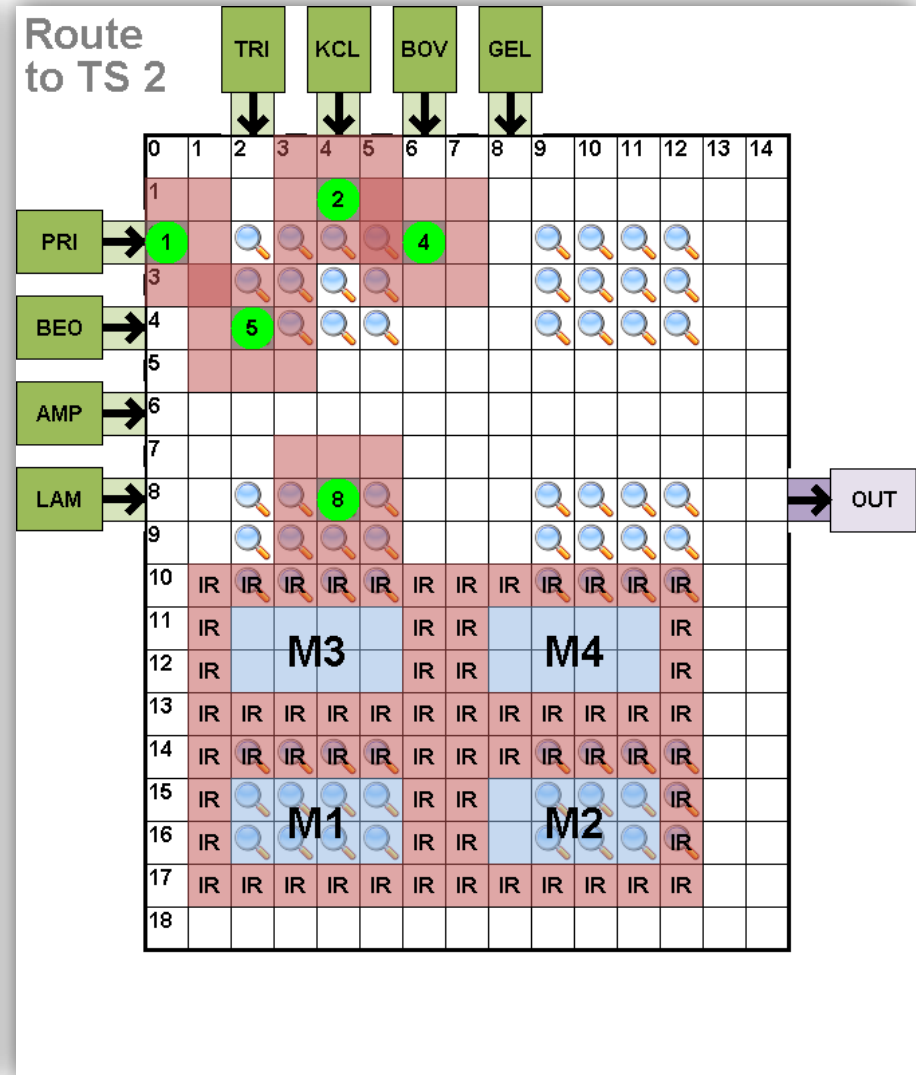


Corresponding Droplet Motion

Compaction Example

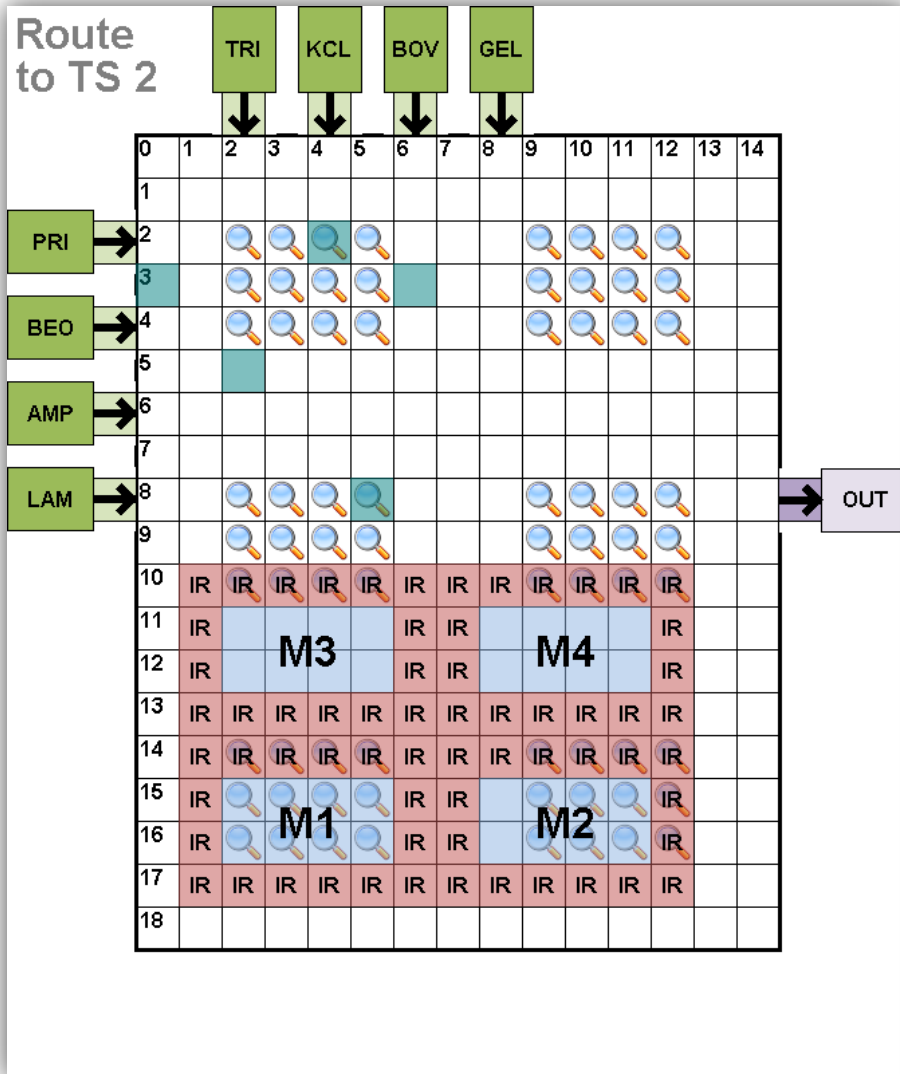


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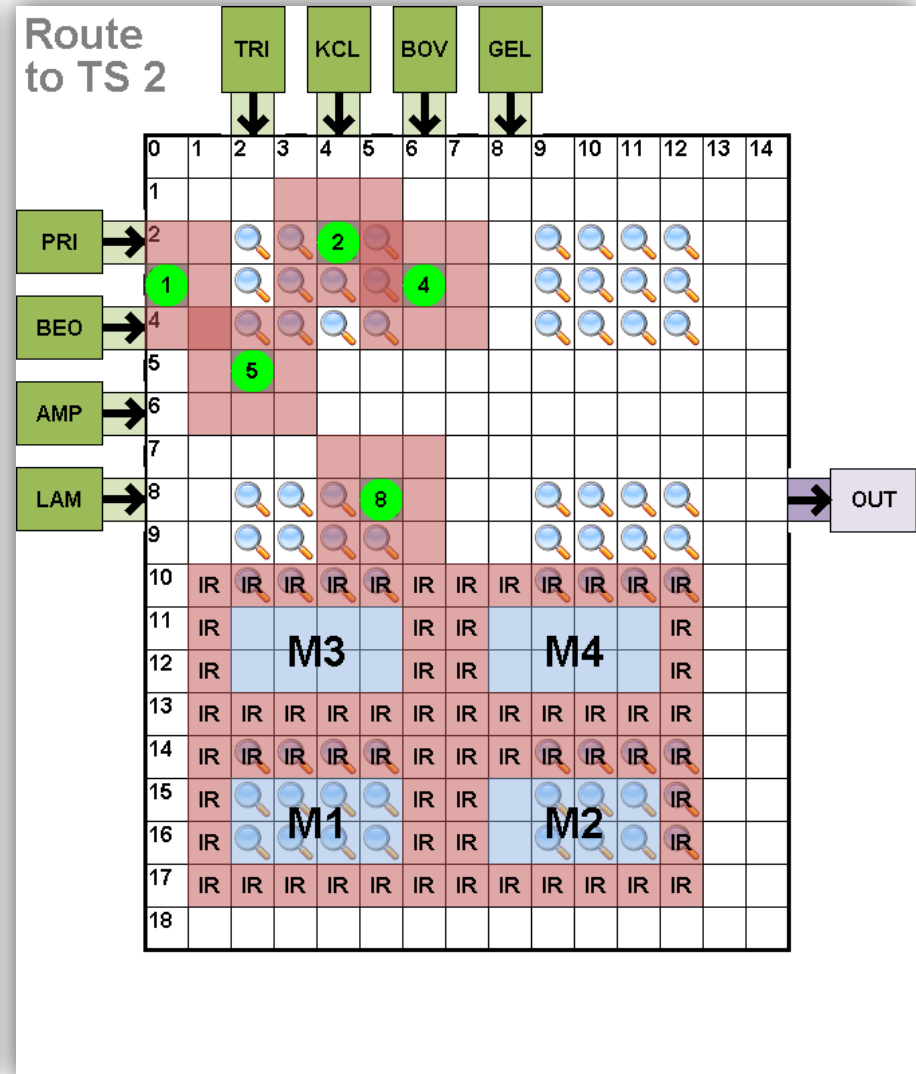


Corresponding Droplet Motion

Compaction Example

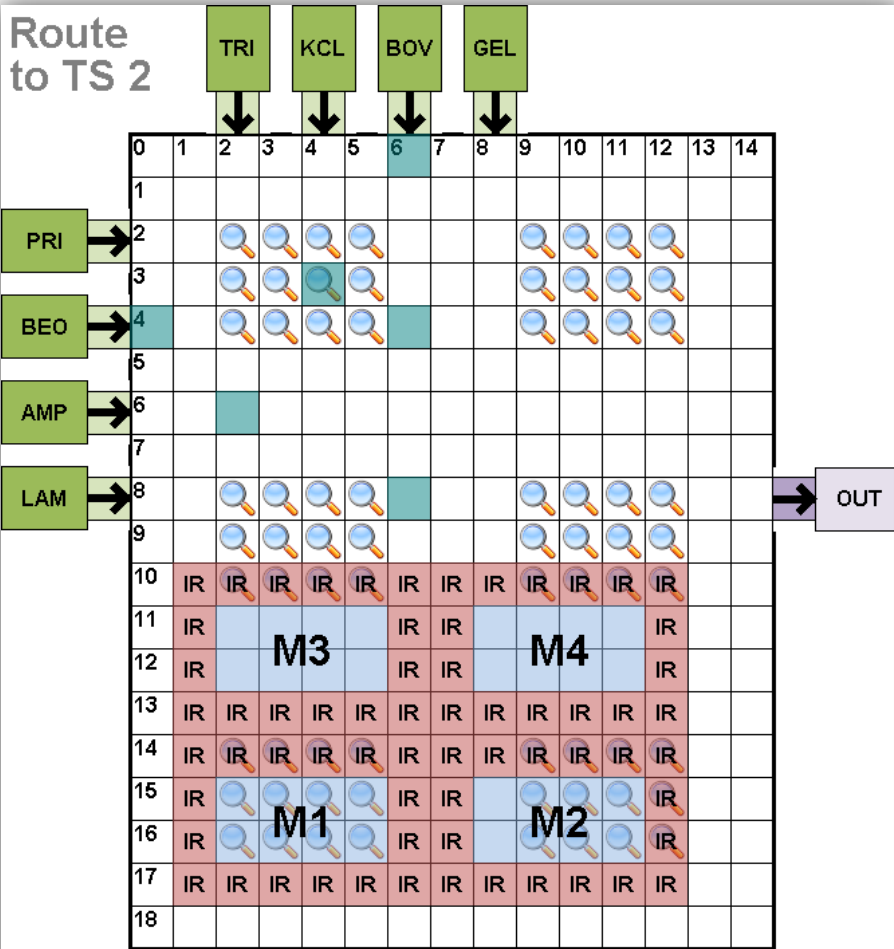


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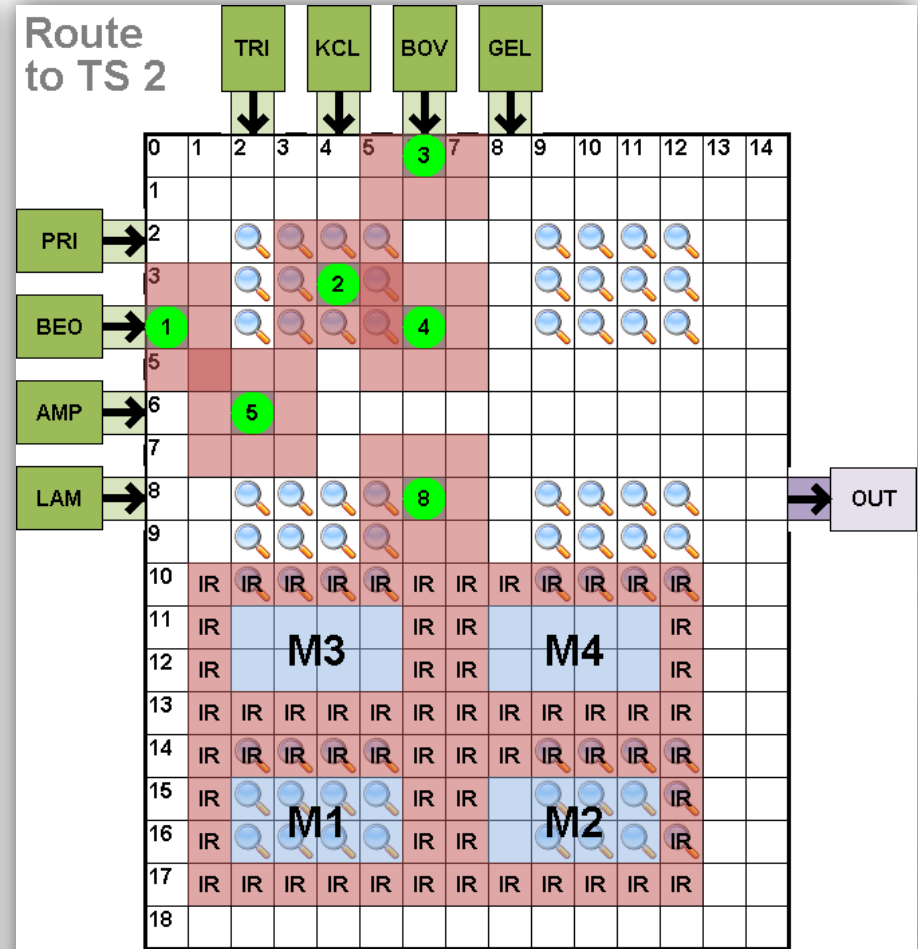


Corresponding Droplet Motion

Compaction Example

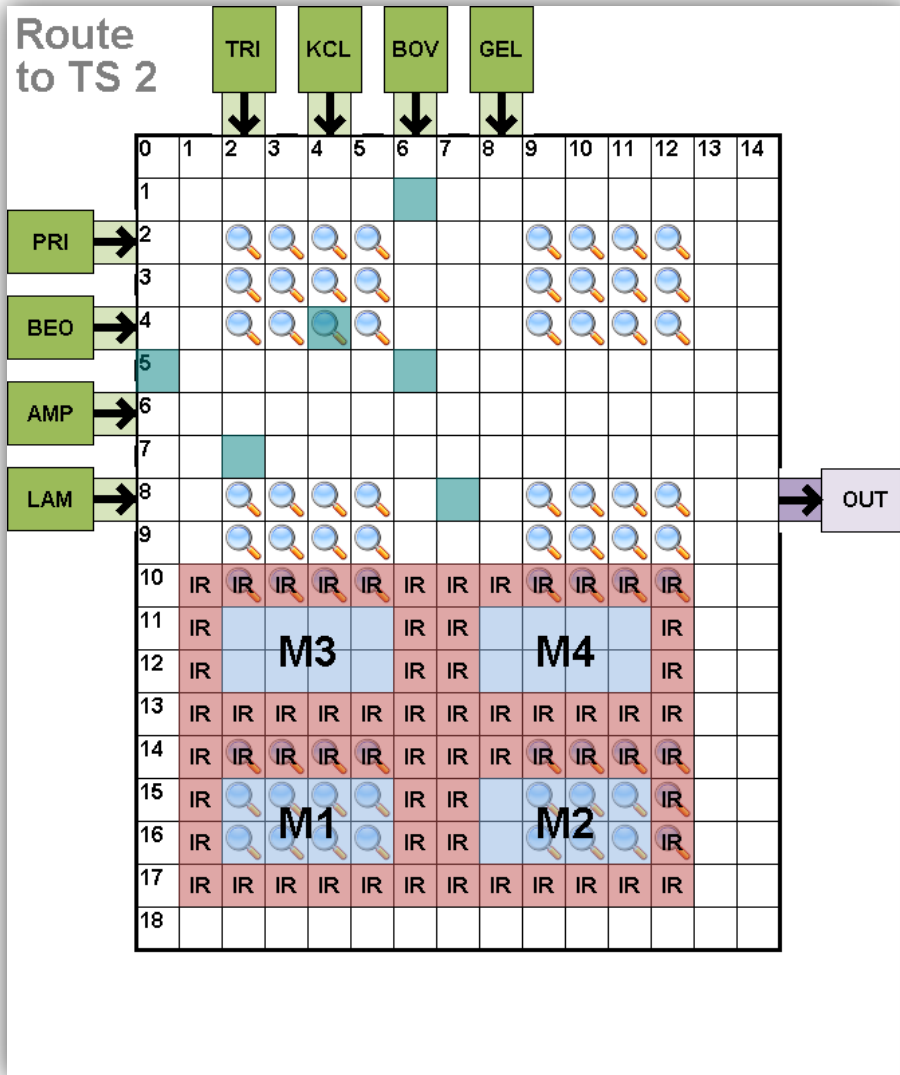


Electrode Activations

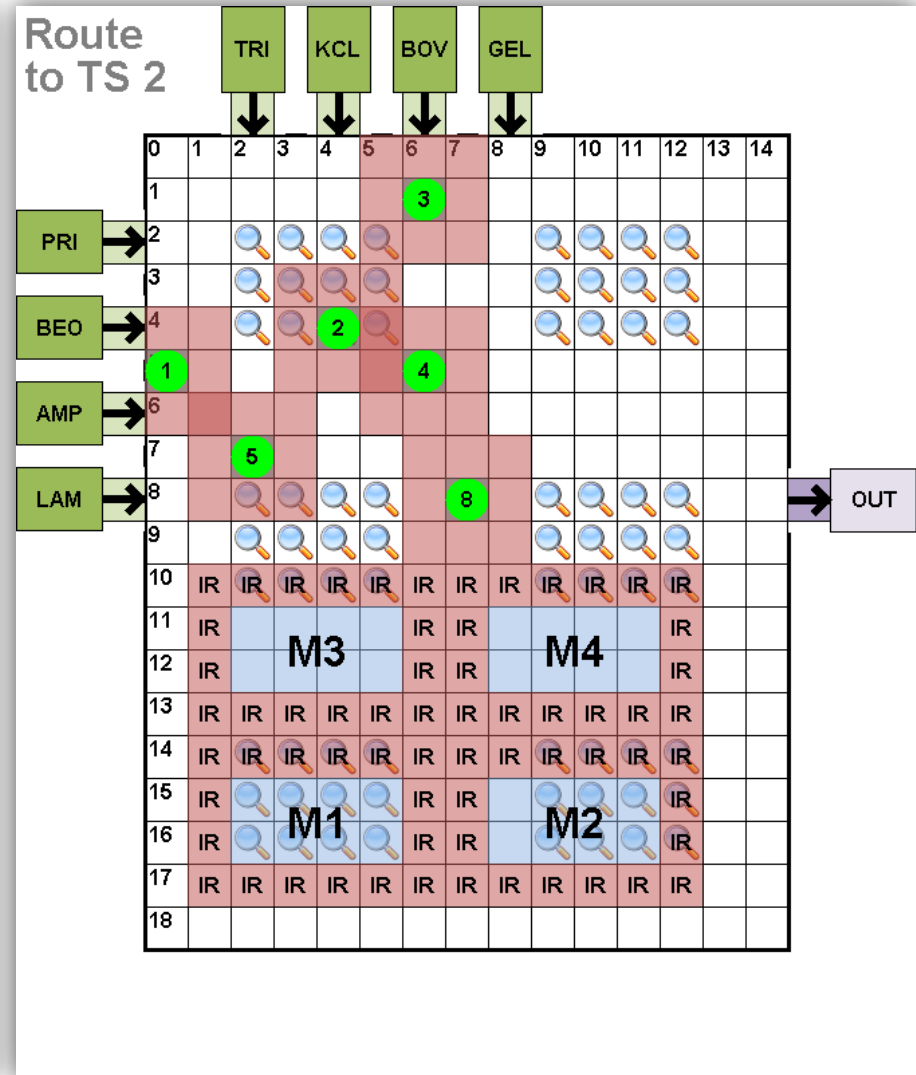


Corresponding Droplet Motion

Compaction Example

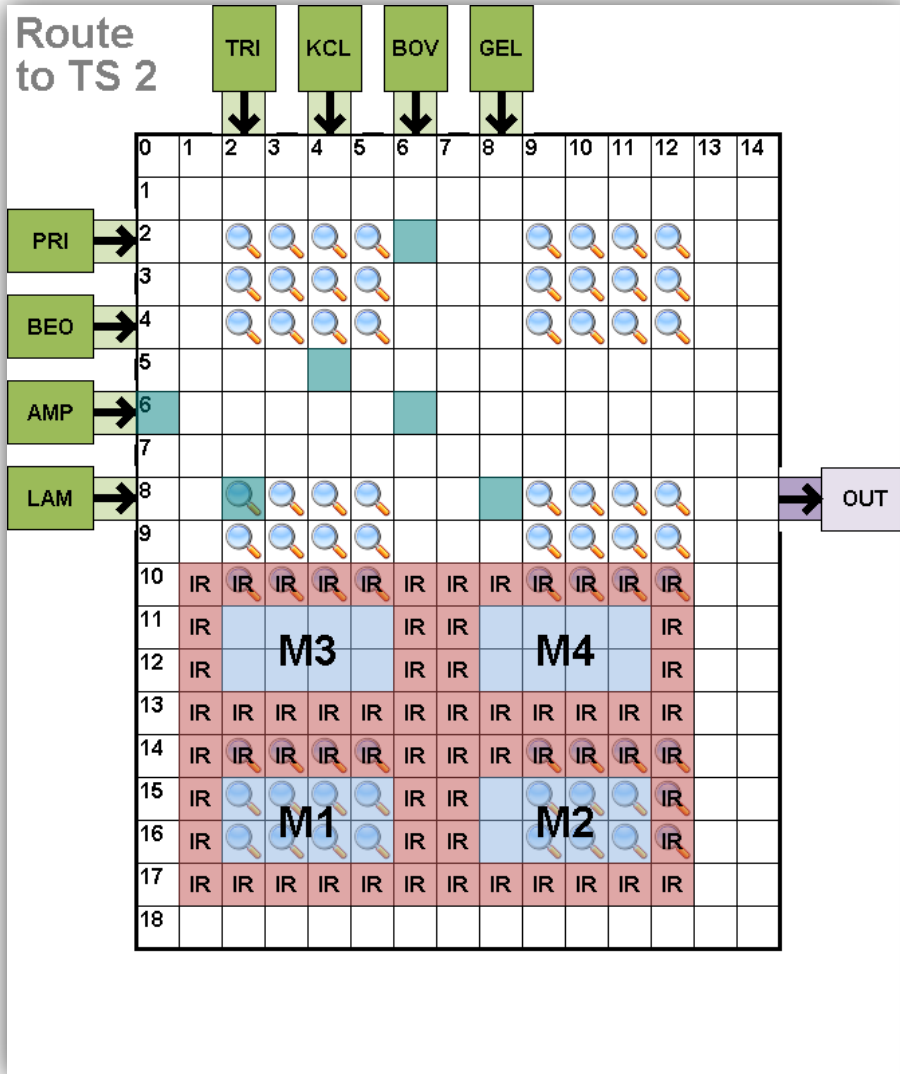


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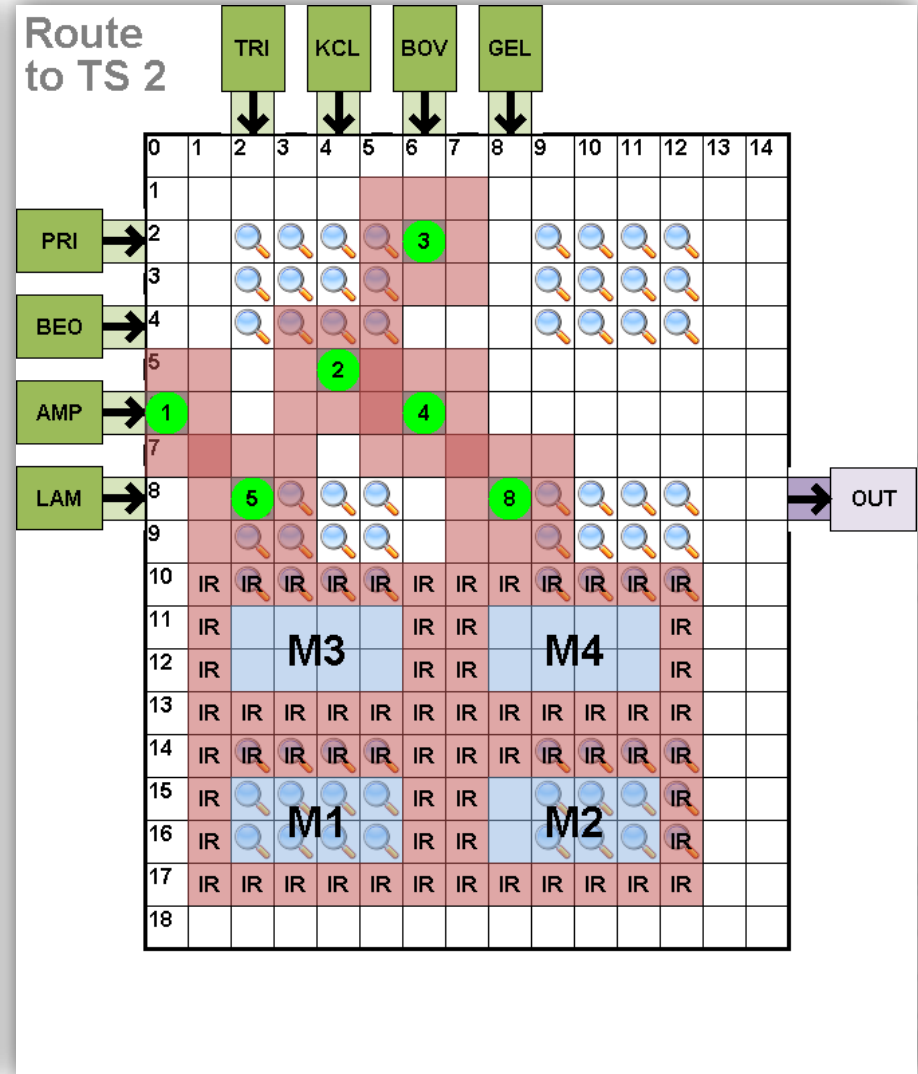


Corresponding Droplet Motion

Compaction Example

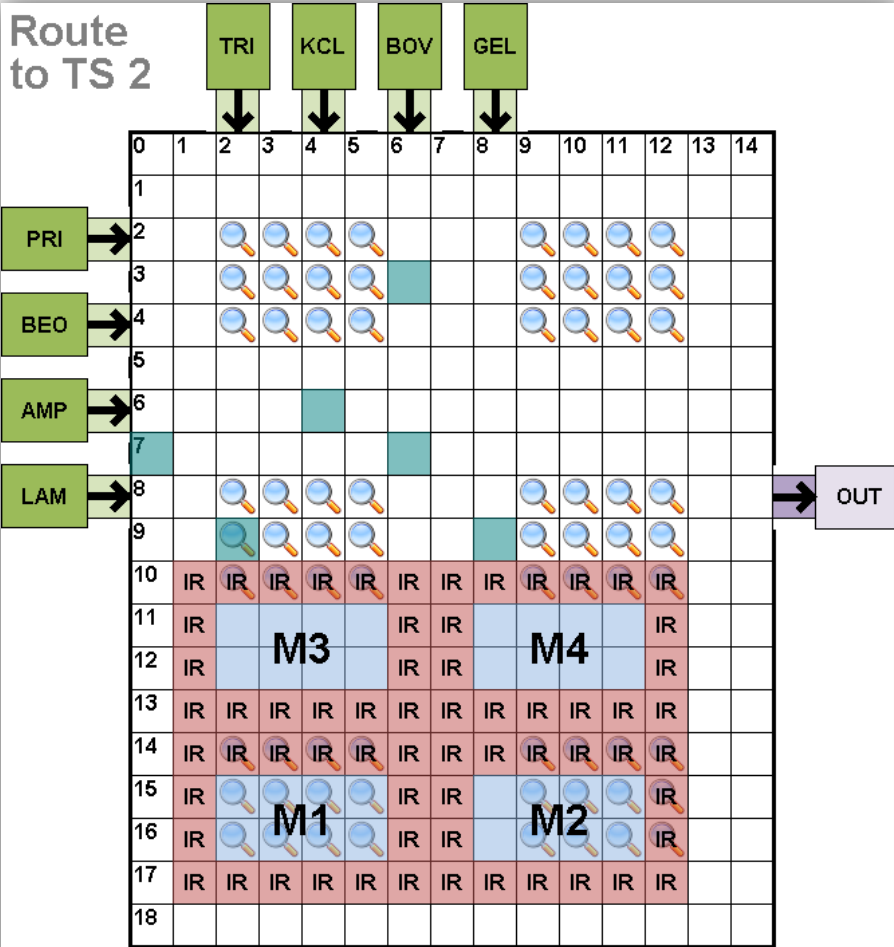


Electrode Activations

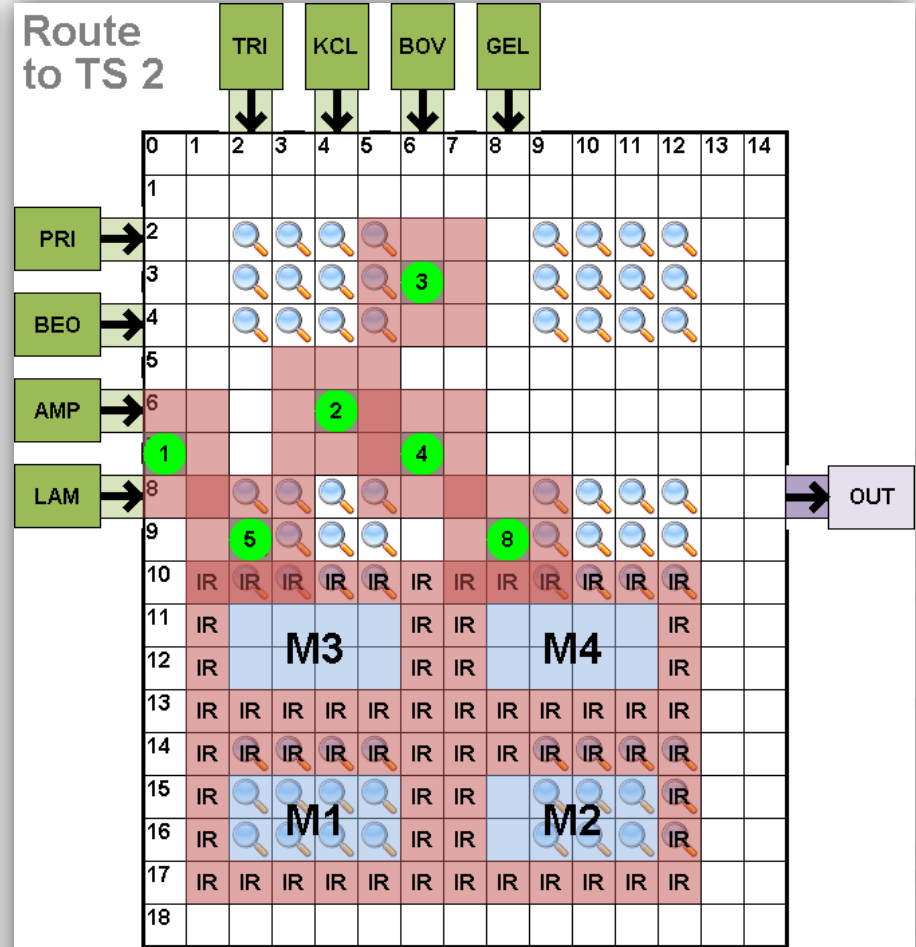


Corresponding Droplet Motion

Compaction Example

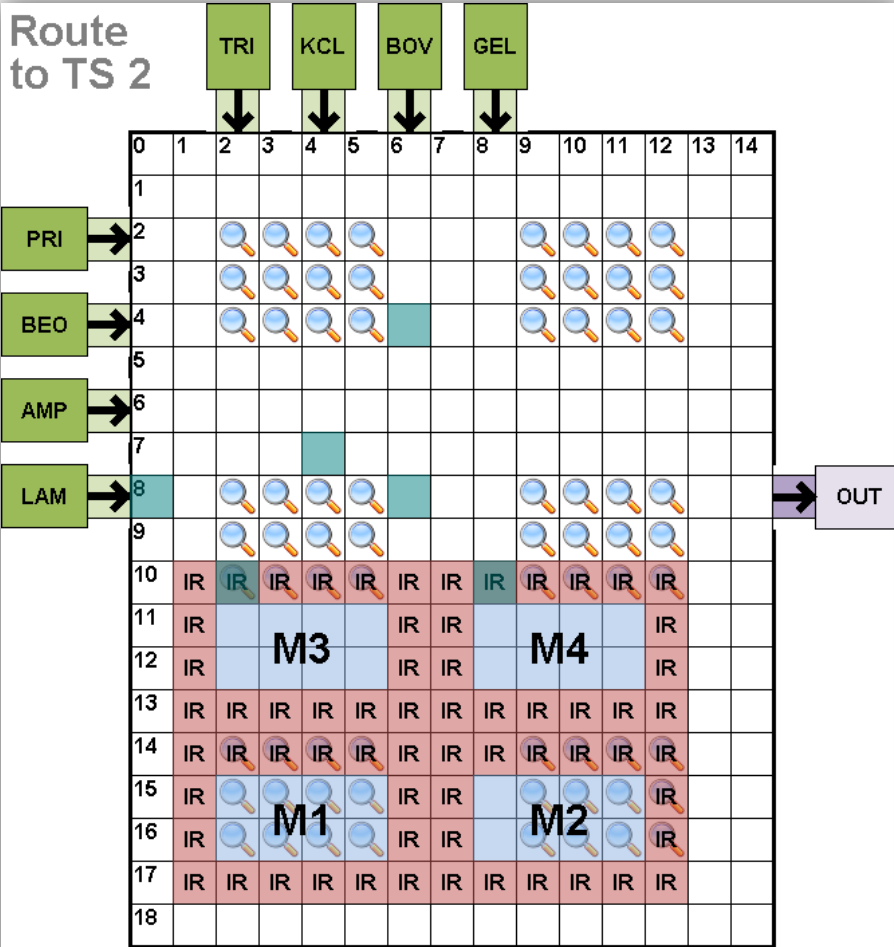


Electrode Activations

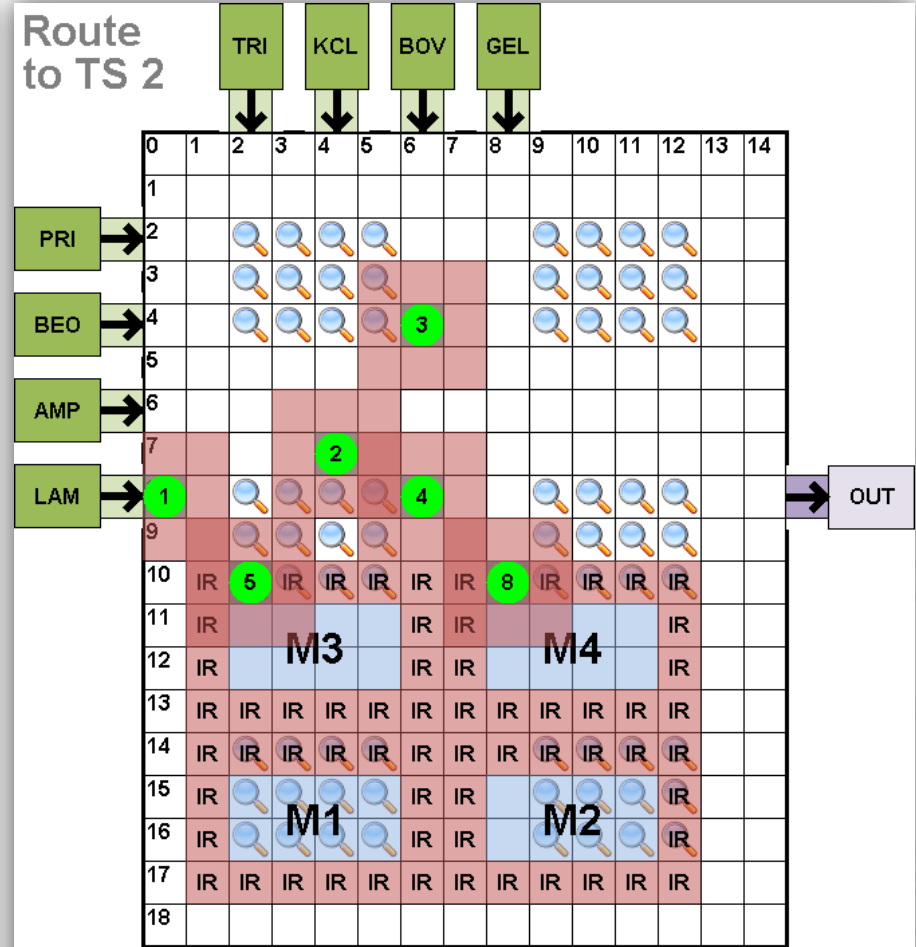


Corresponding Droplet Motion

Compaction Example

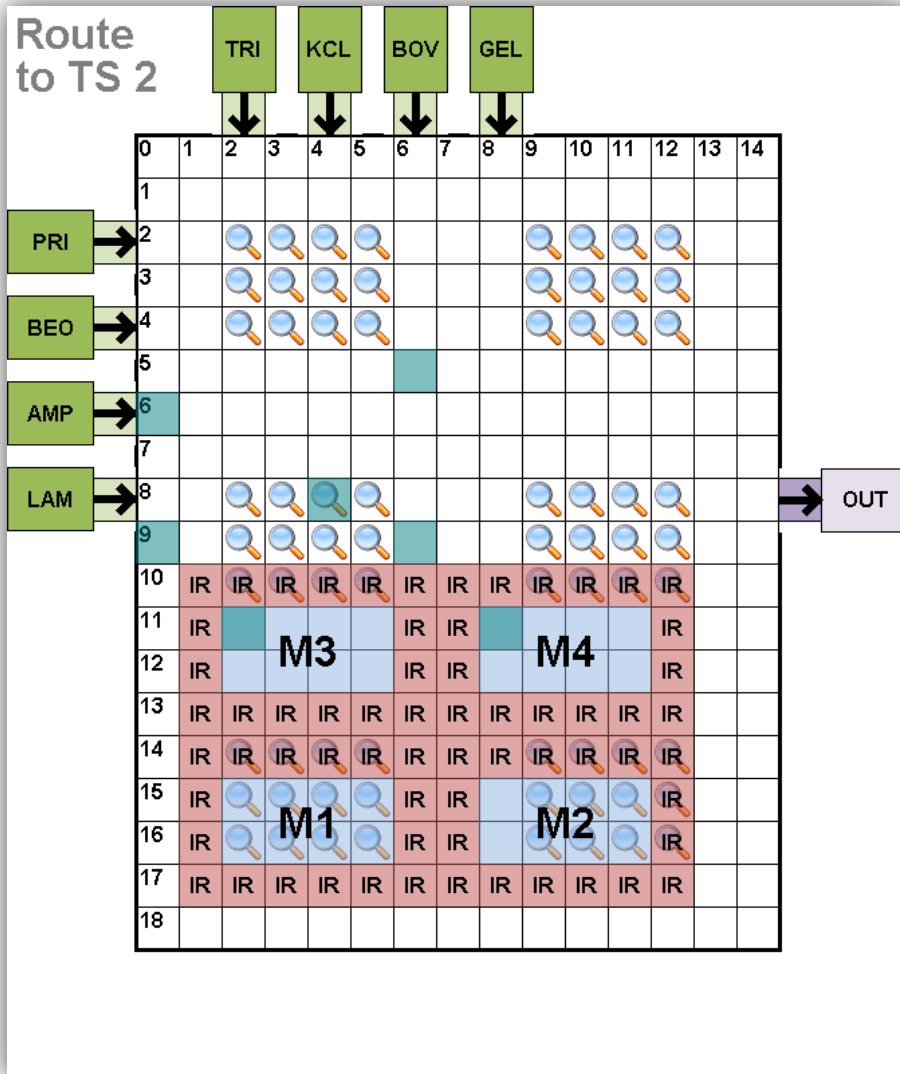


Electrode Activations

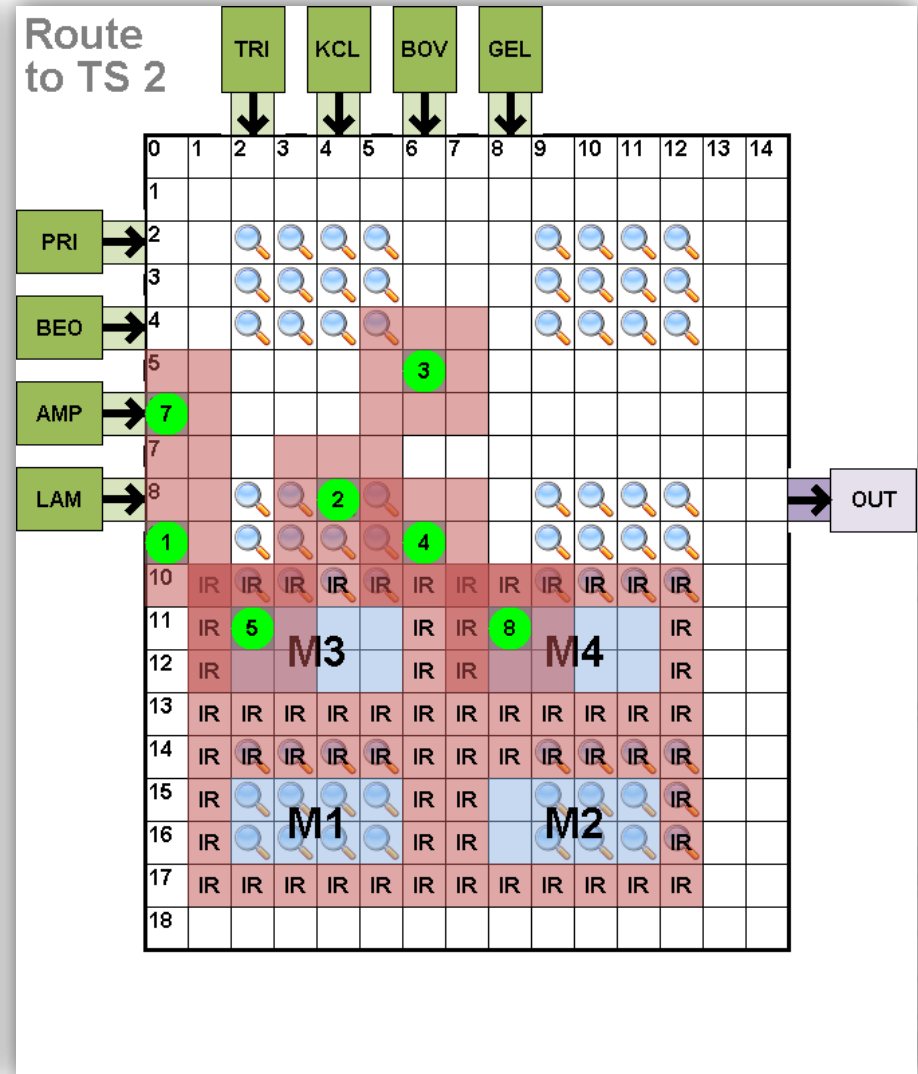


Corresponding Droplet Motion

Compaction Example

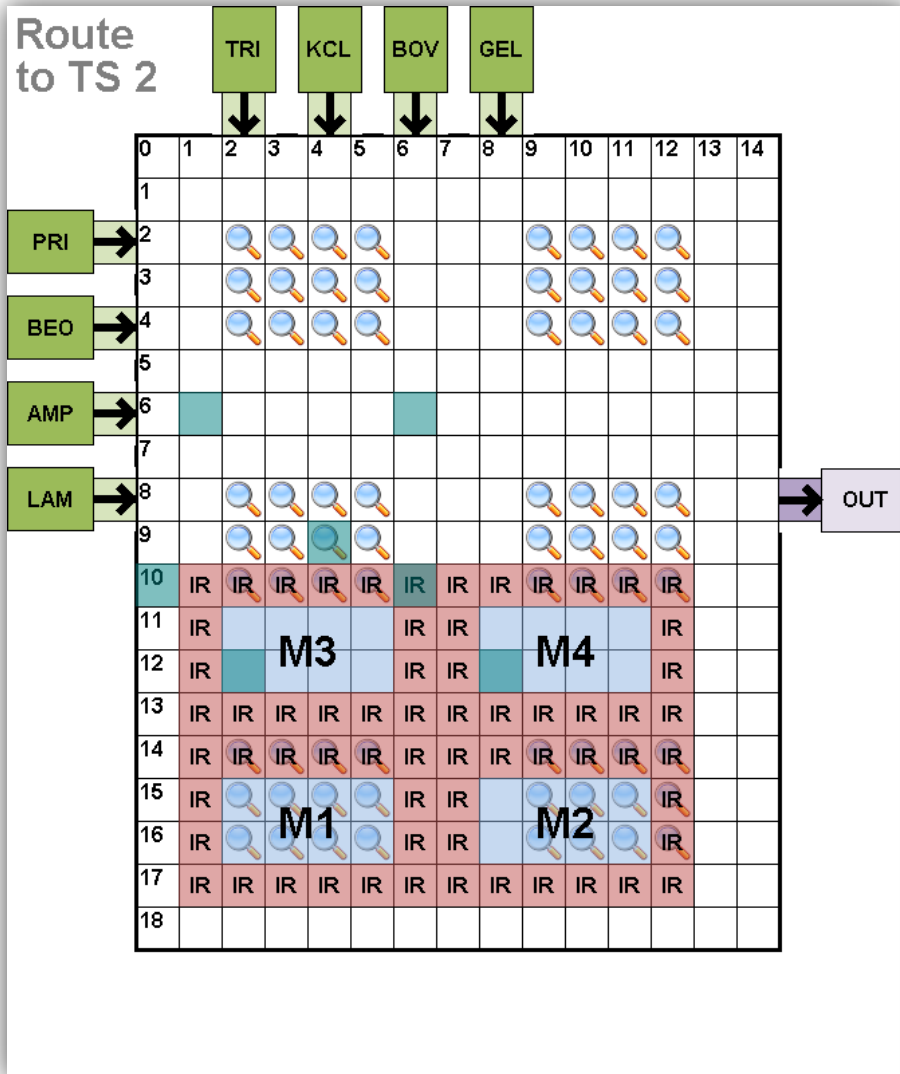


Electrode Activations

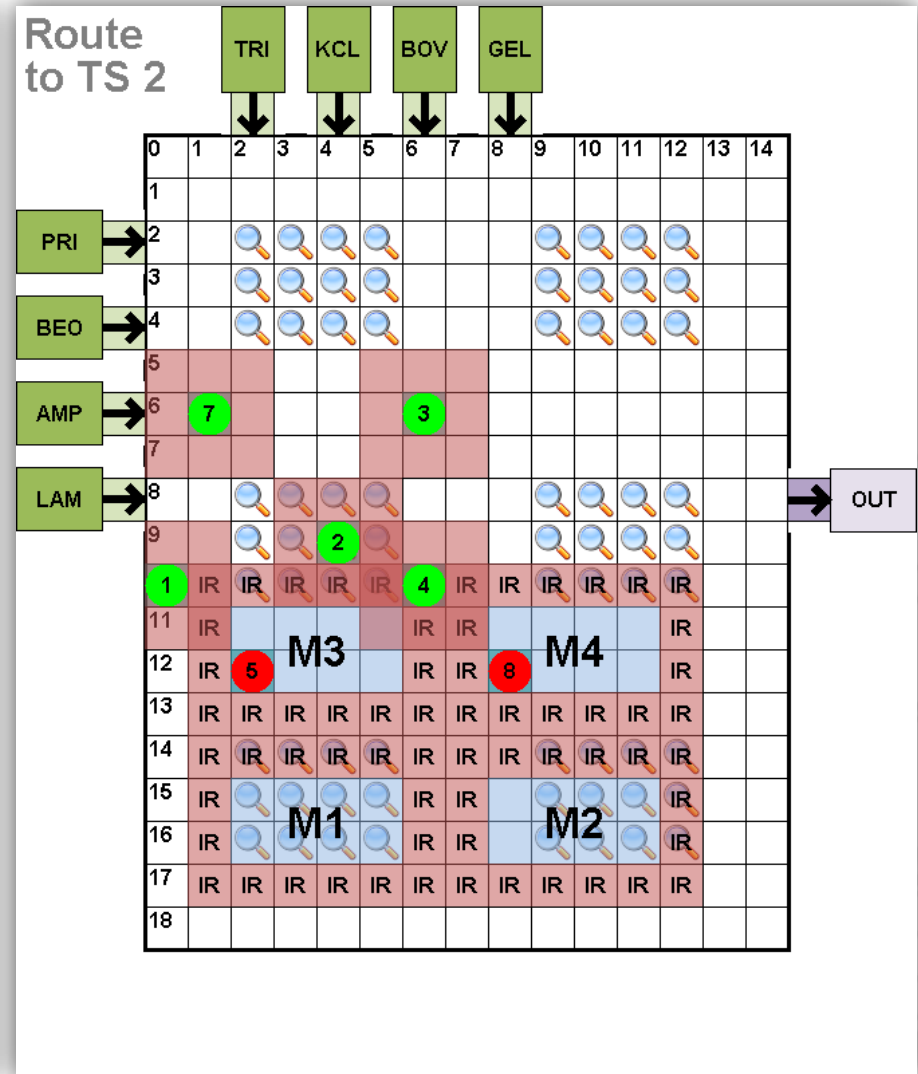


Corresponding Droplet Motion

Compaction Example

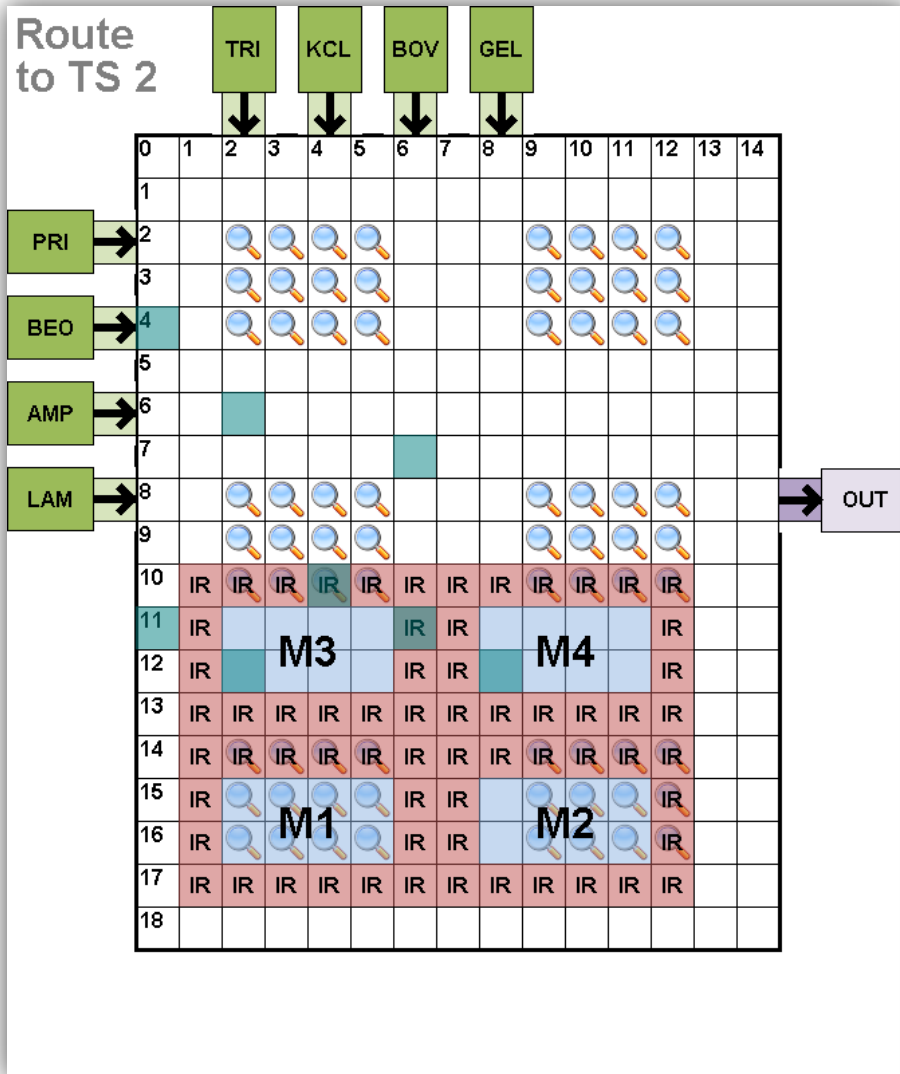


Electrode Activations

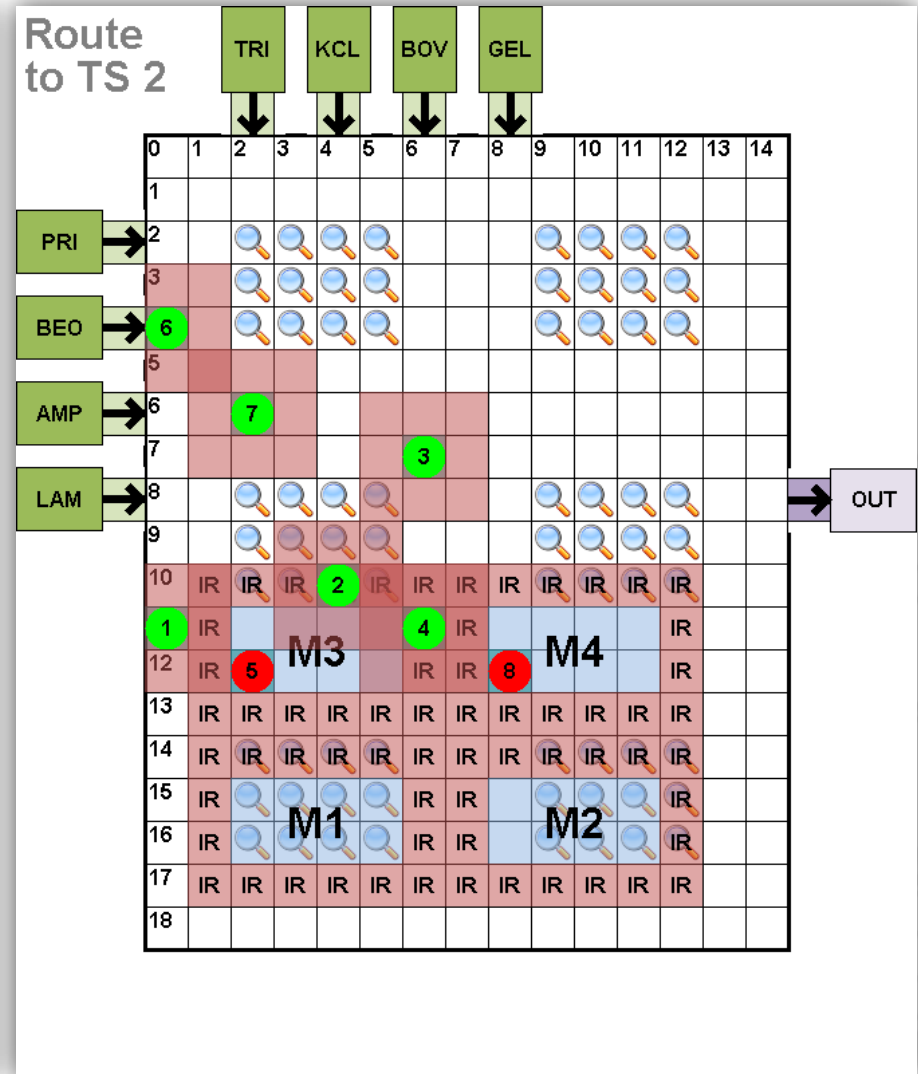


Corresponding Droplet Motion

Compaction Example

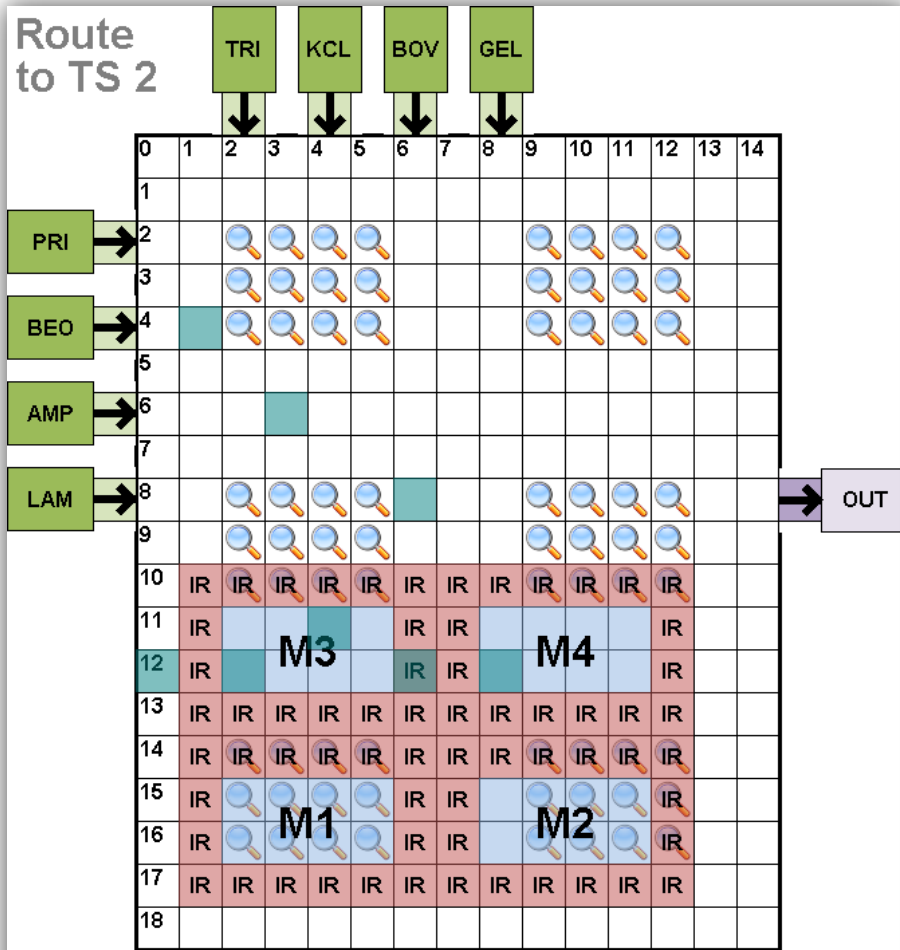


Electrode Activations

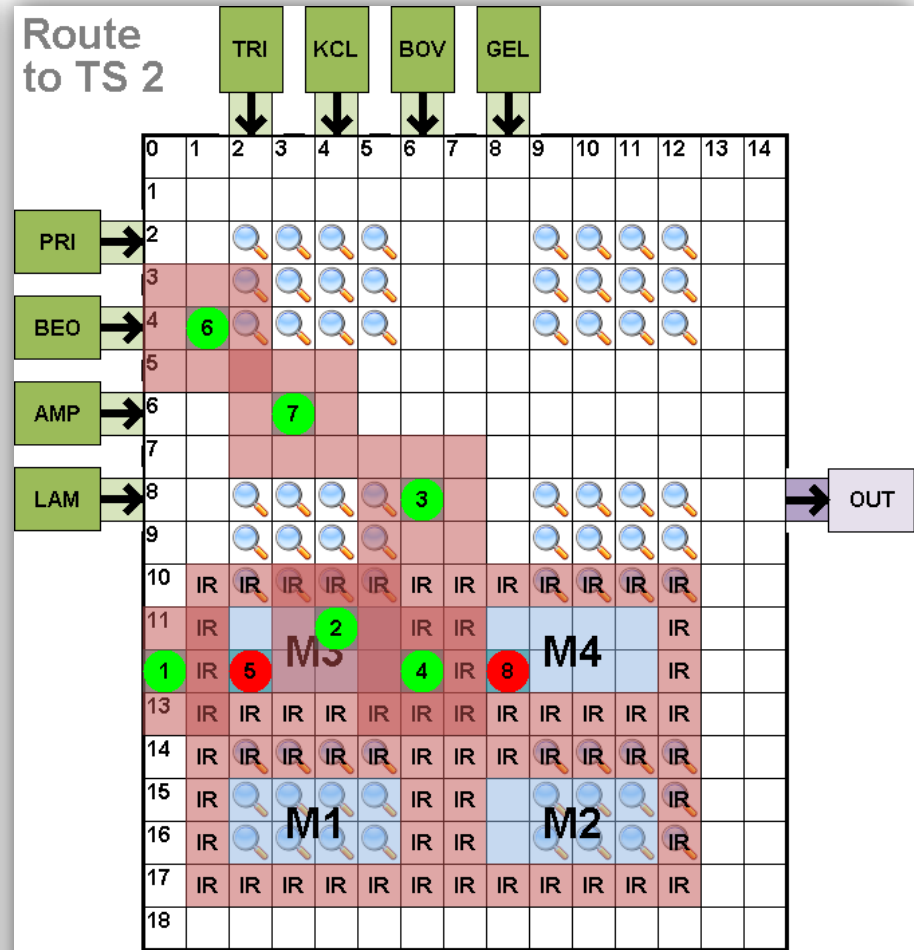


Corresponding Droplet Motion

Compaction Example

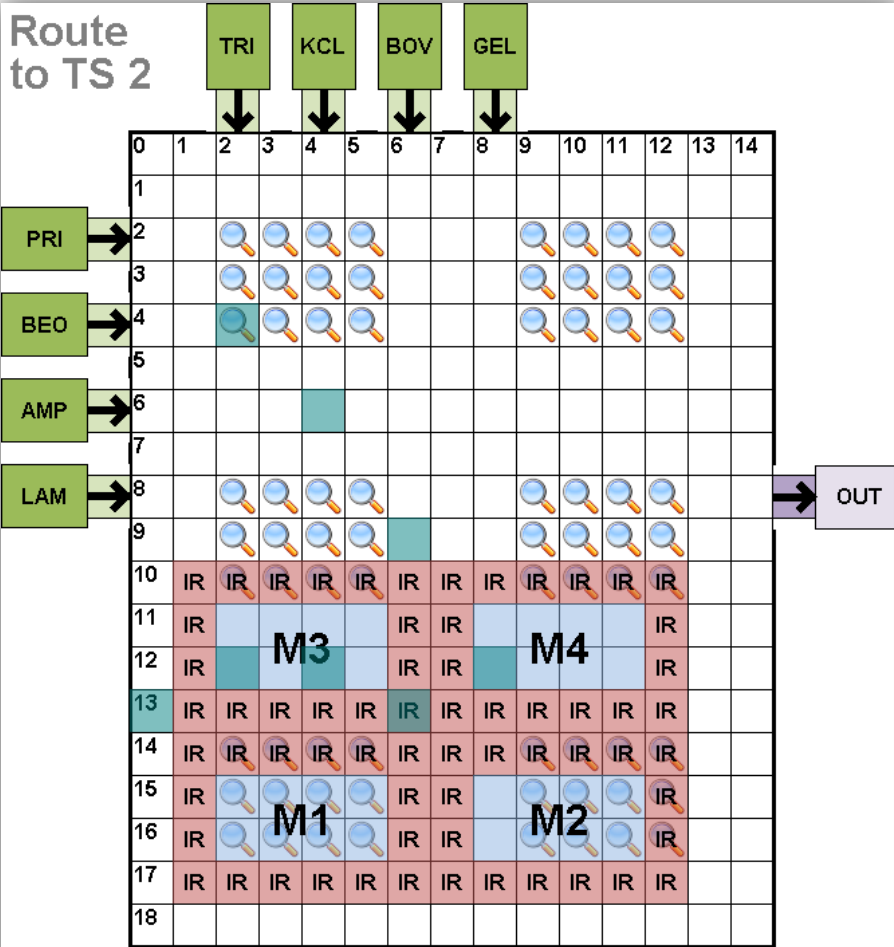


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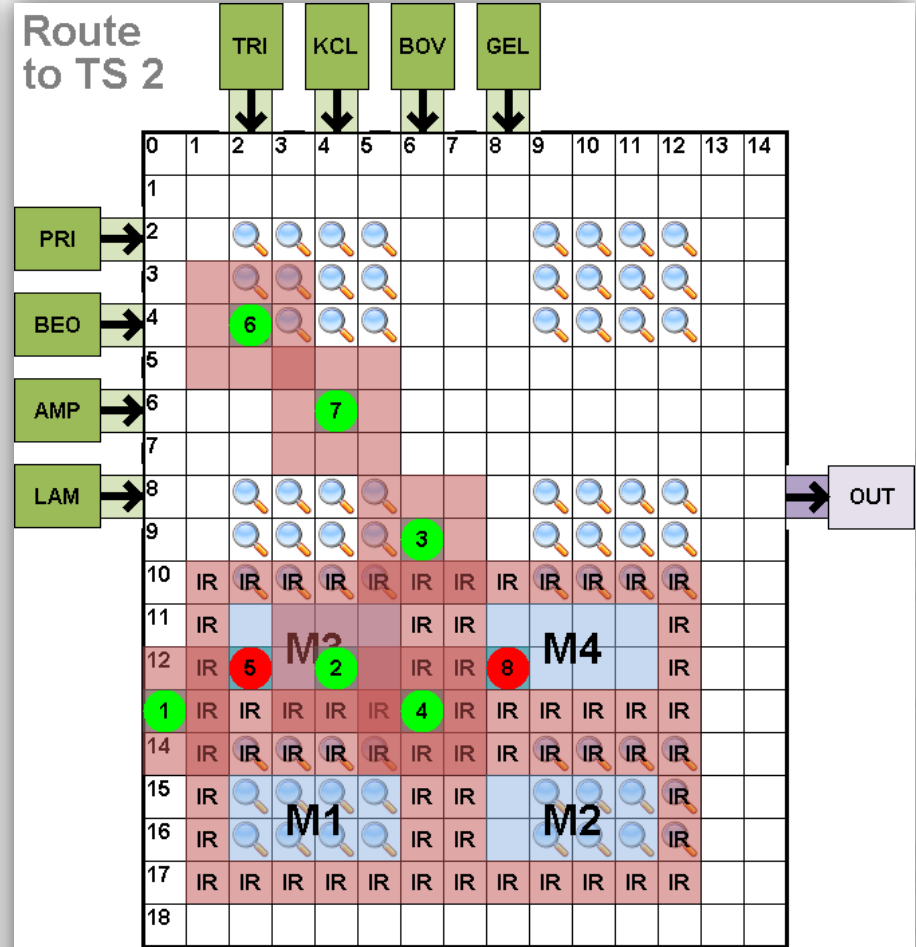


Corresponding Droplet Motion

Compaction Example

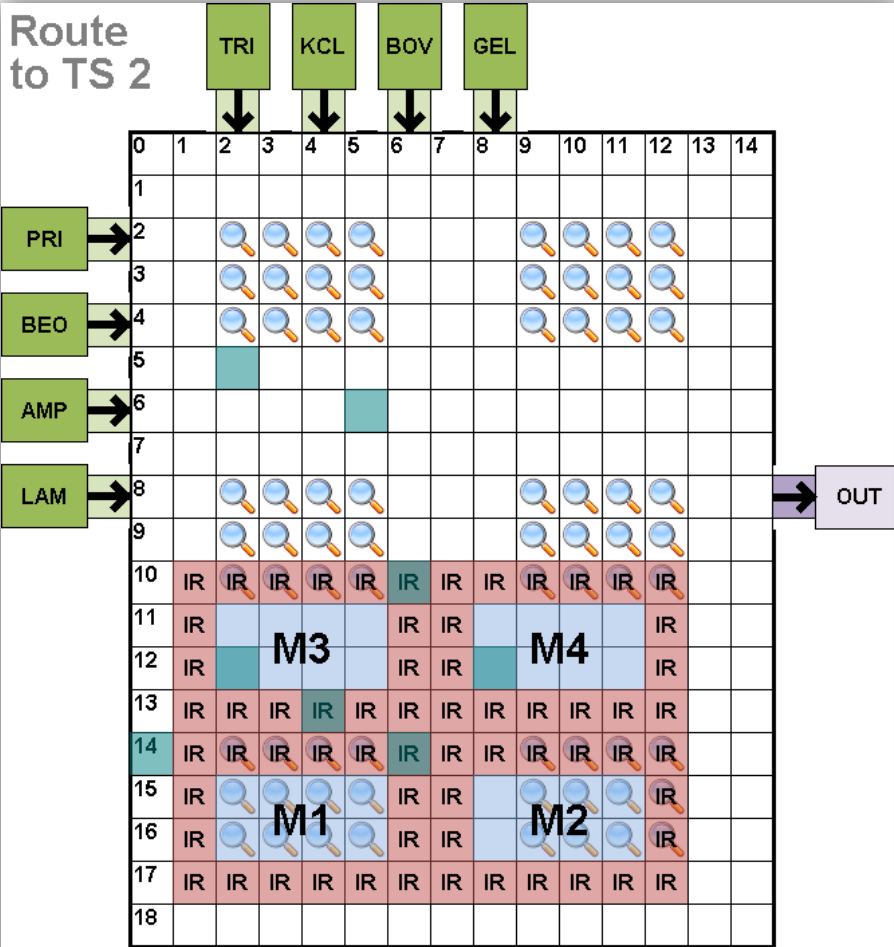


Electrode Activations

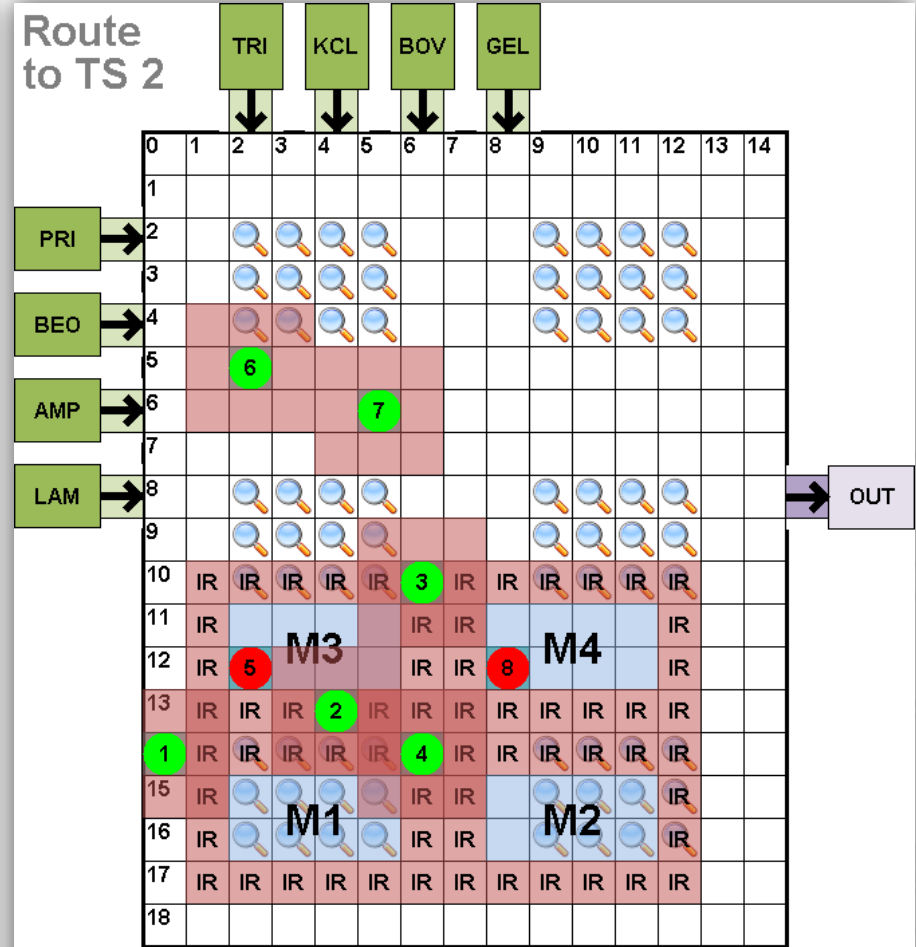


Corresponding Droplet Motion

Compaction Example

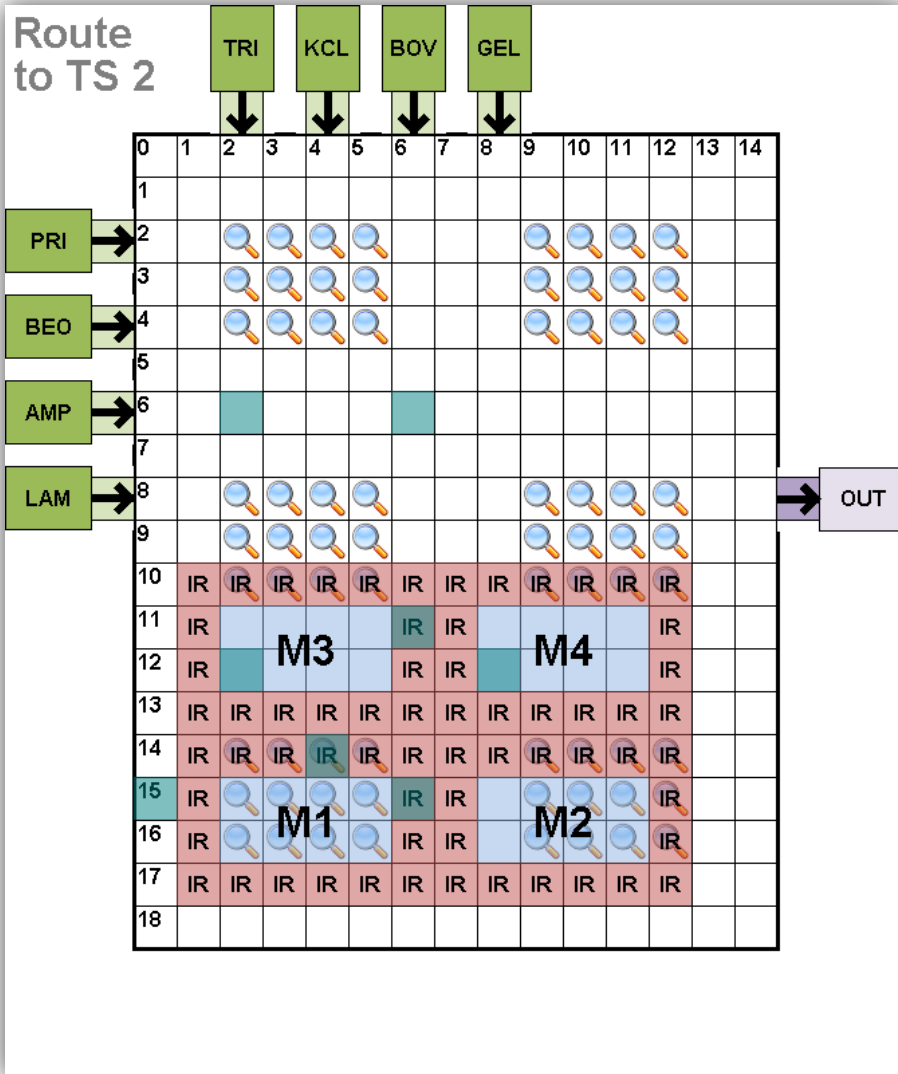


Electrode Activations

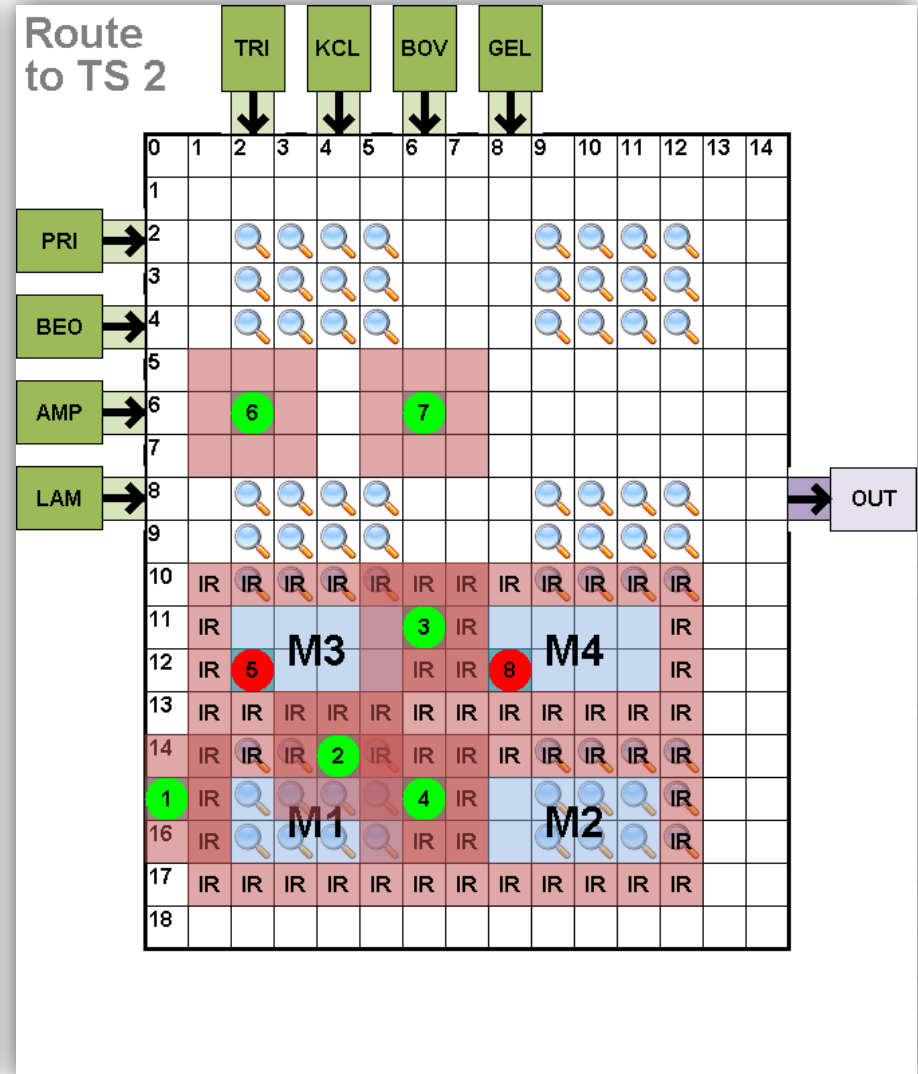


Corresponding Droplet Motion

Compaction Example

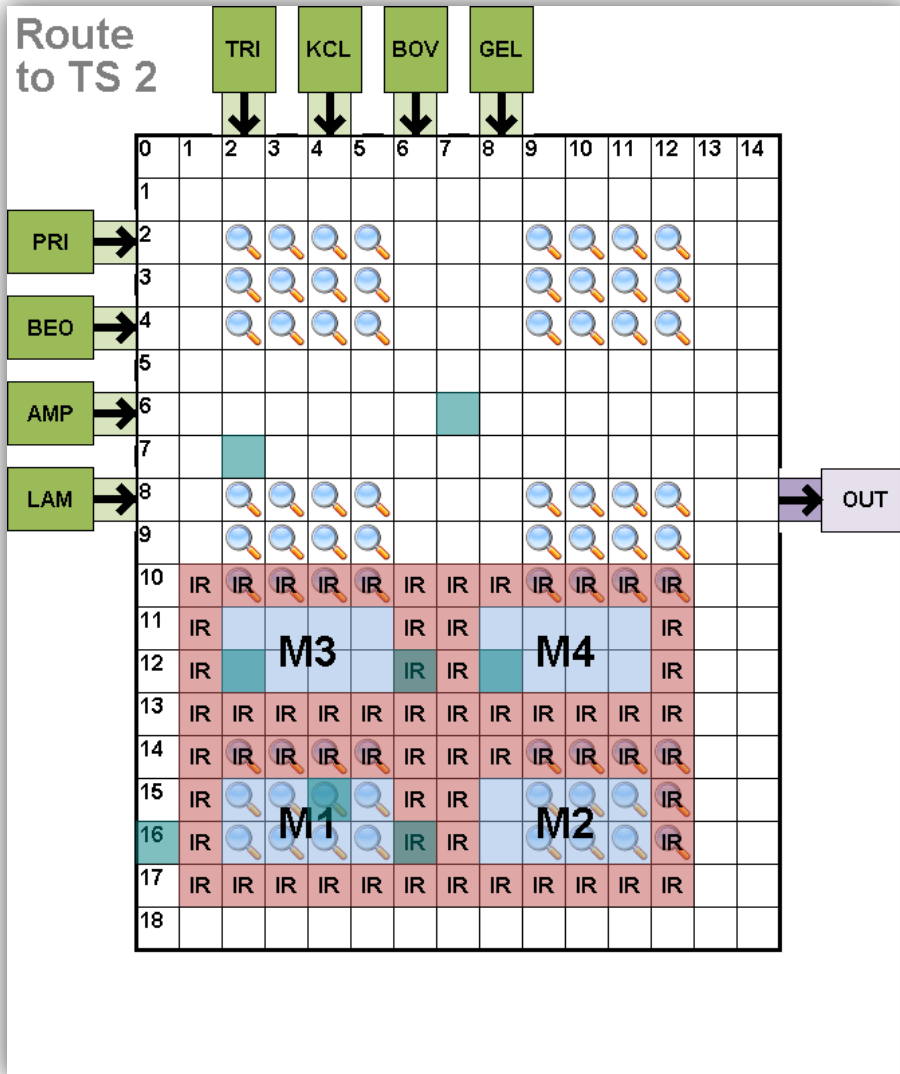


Electrode Activations

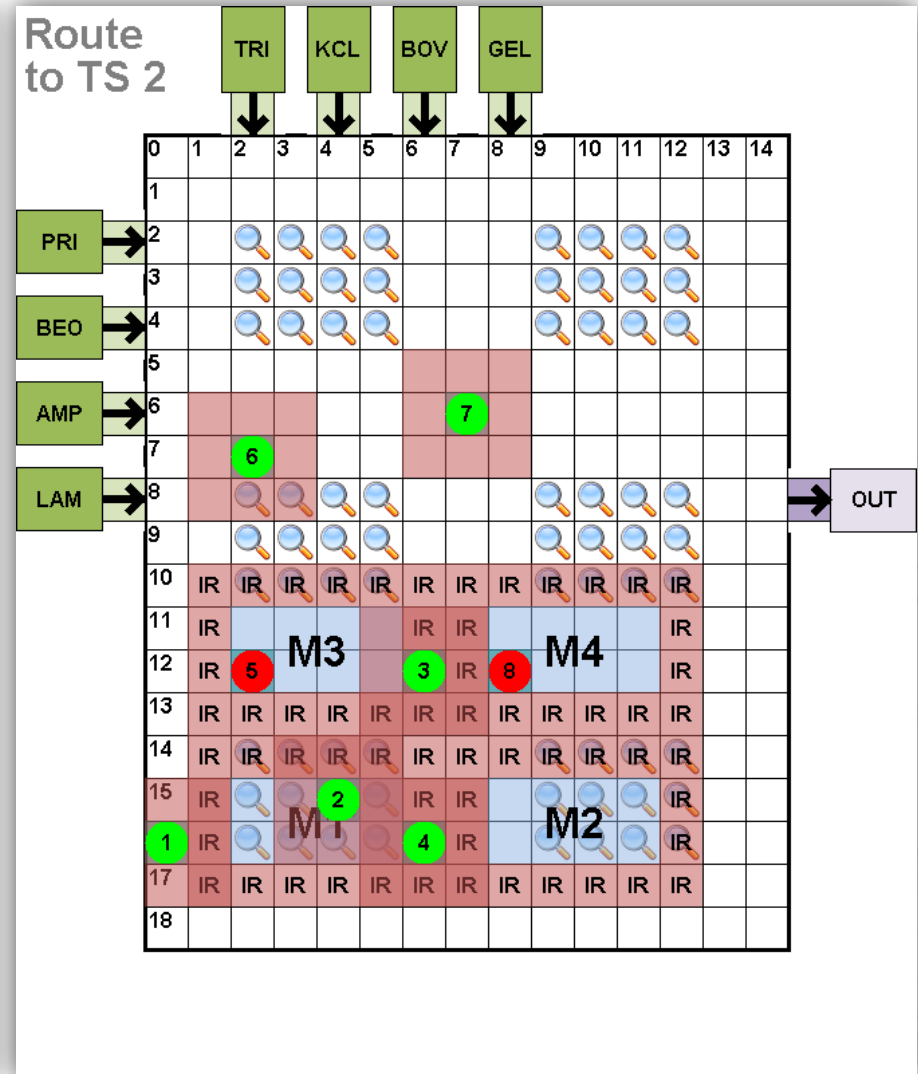


Corresponding Droplet Motion

Compaction Example

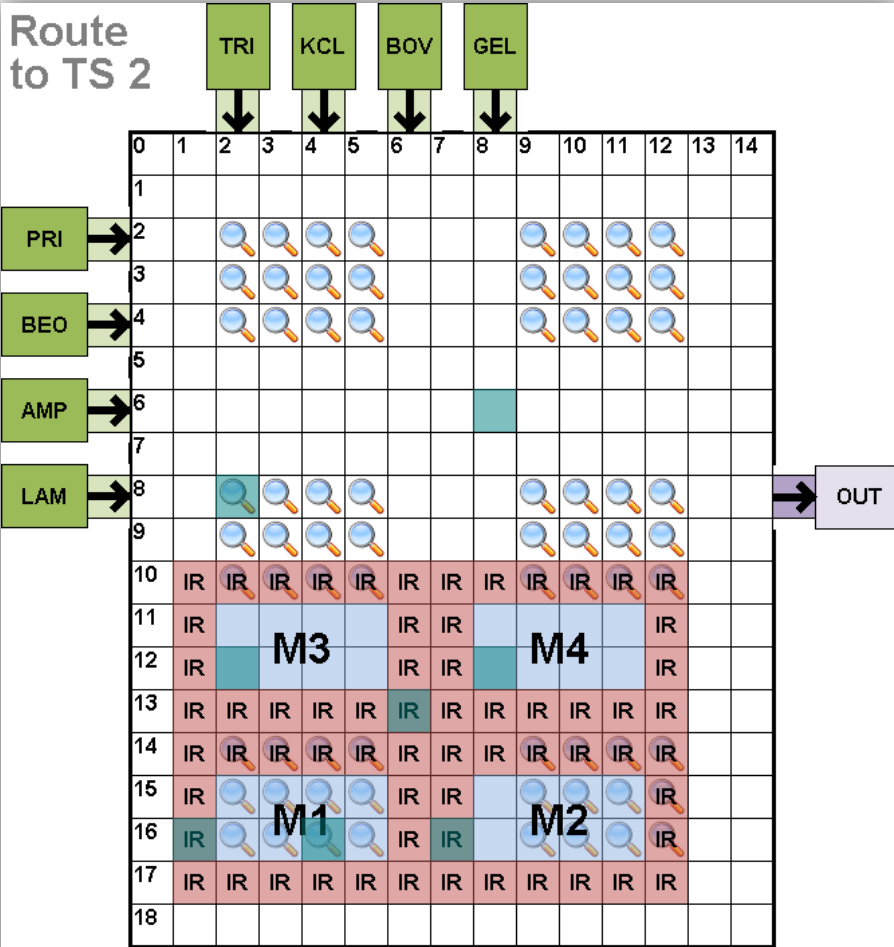


Electrode Activations

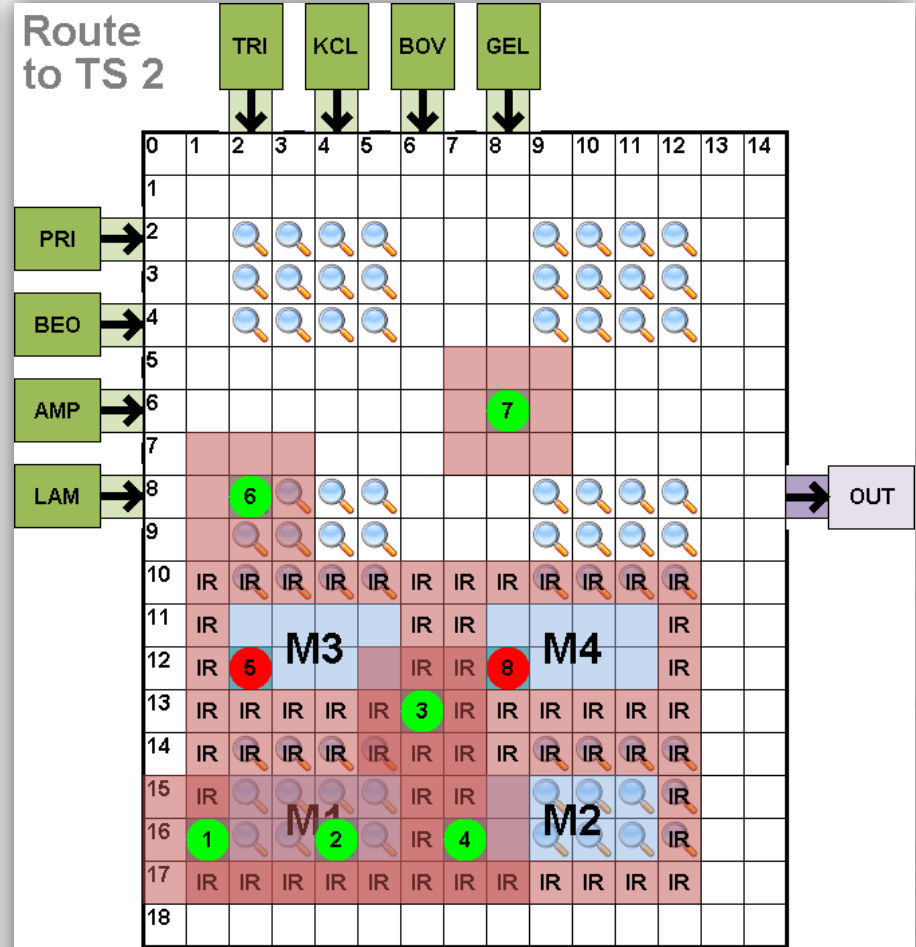


Corresponding Droplet Motion

Compaction Example

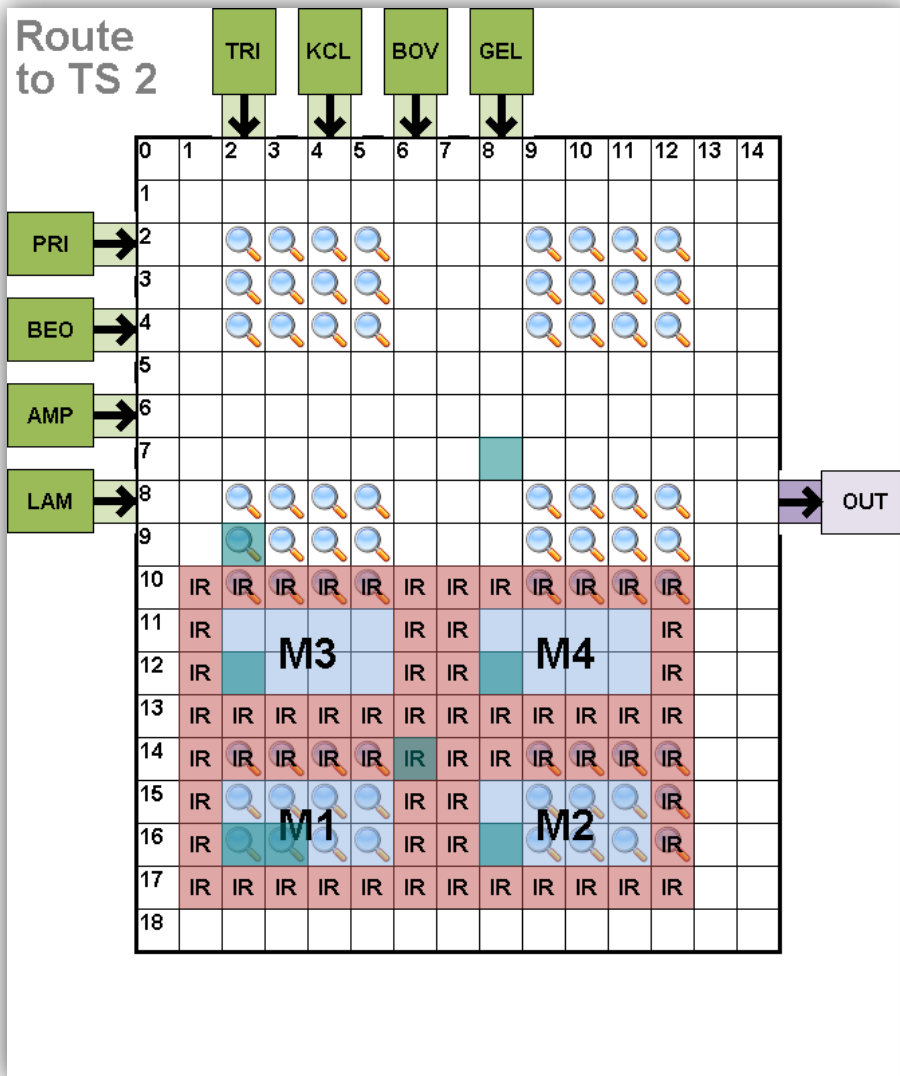


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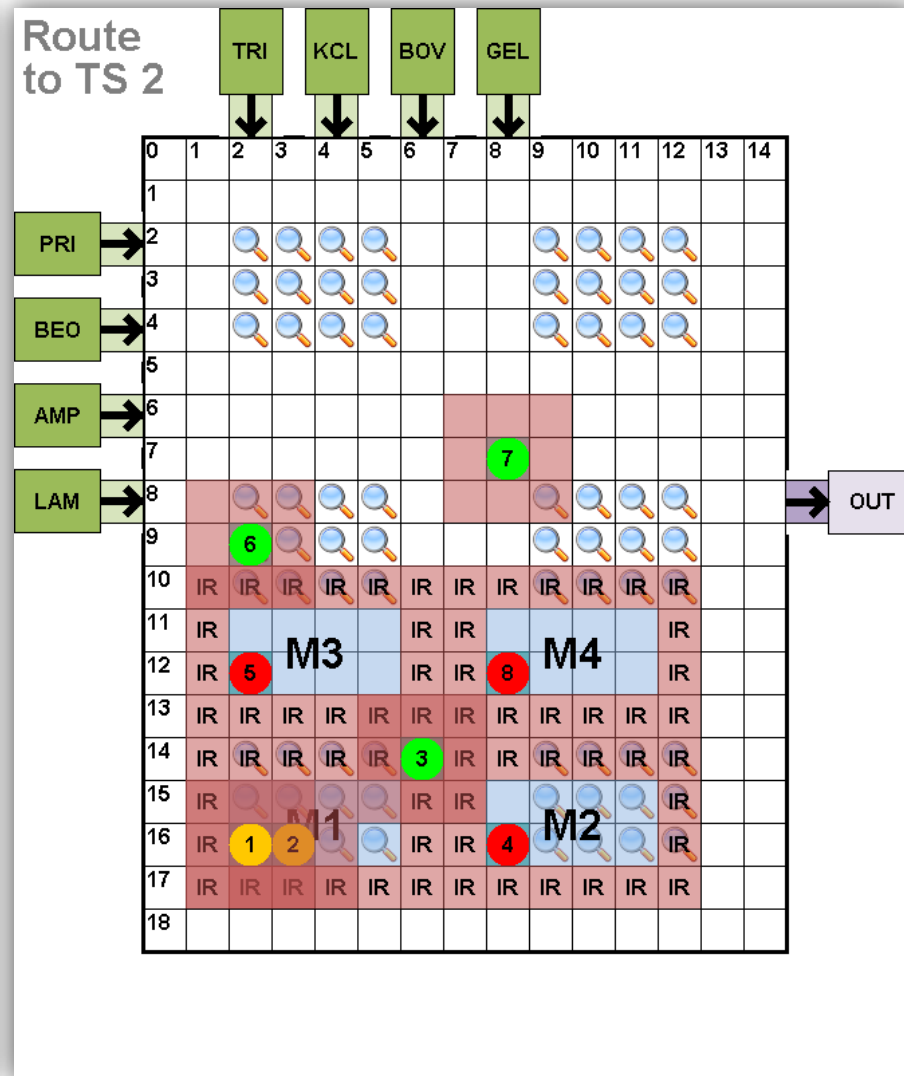


Corresponding Droplet Motion

Compaction Example

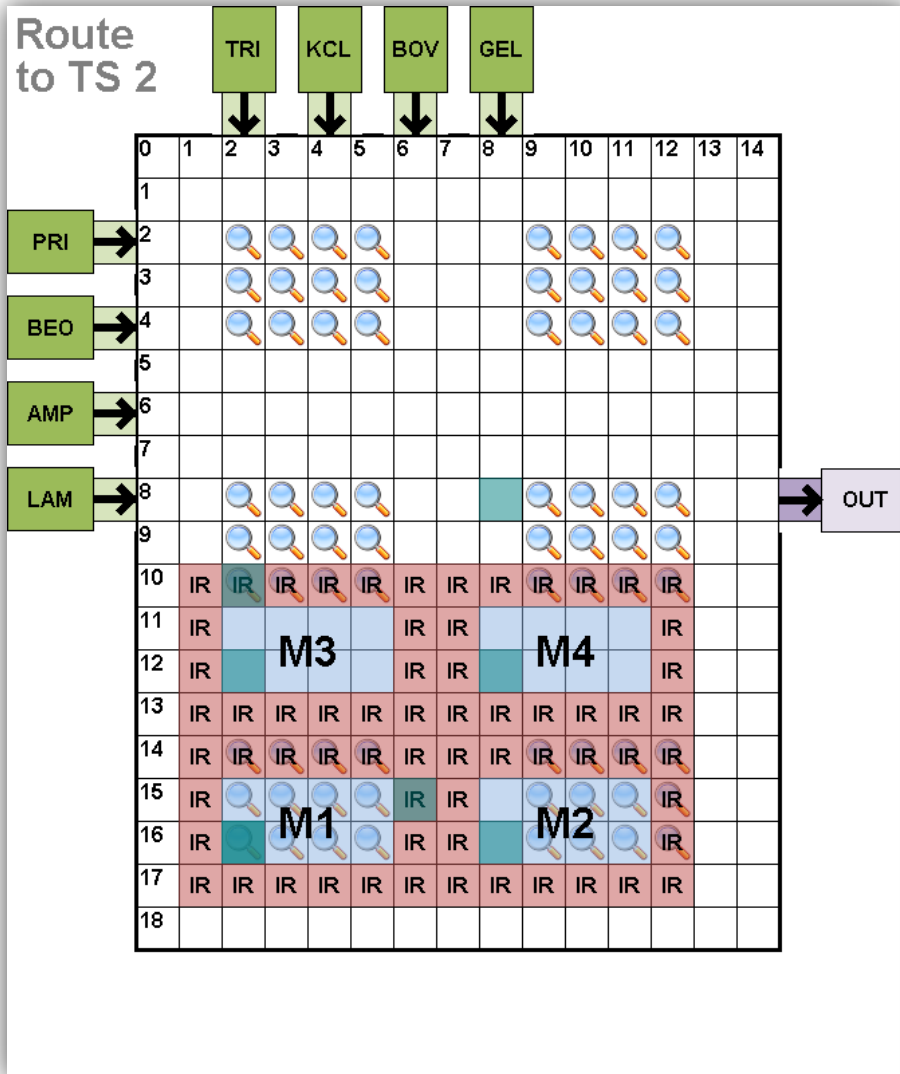


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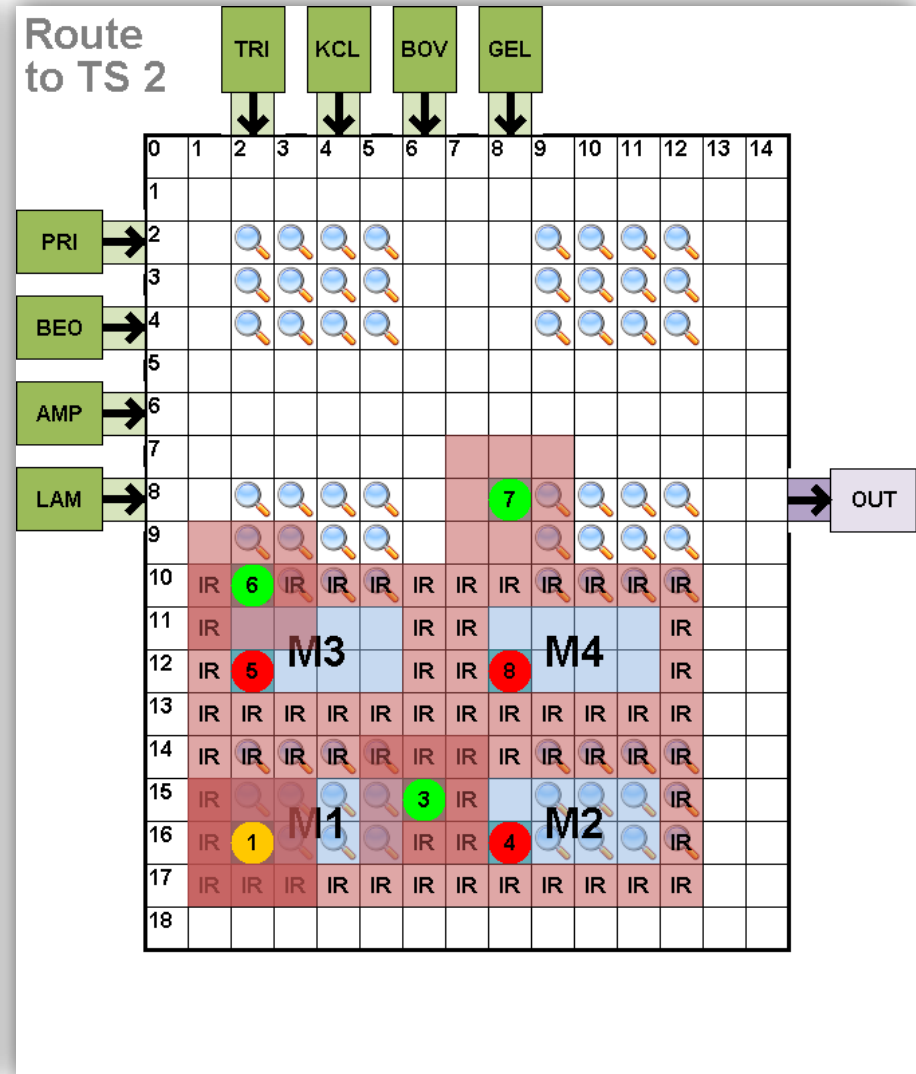


Corresponding Droplet Motion

Compaction Example

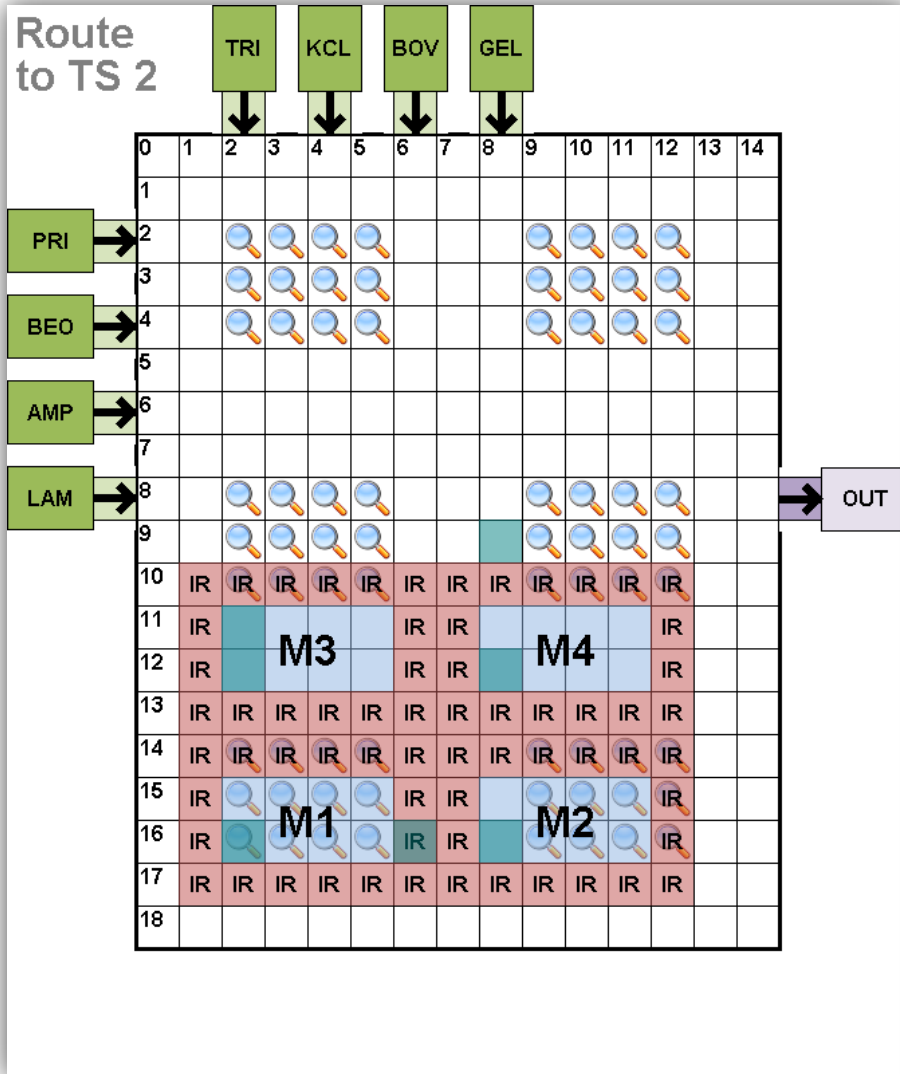


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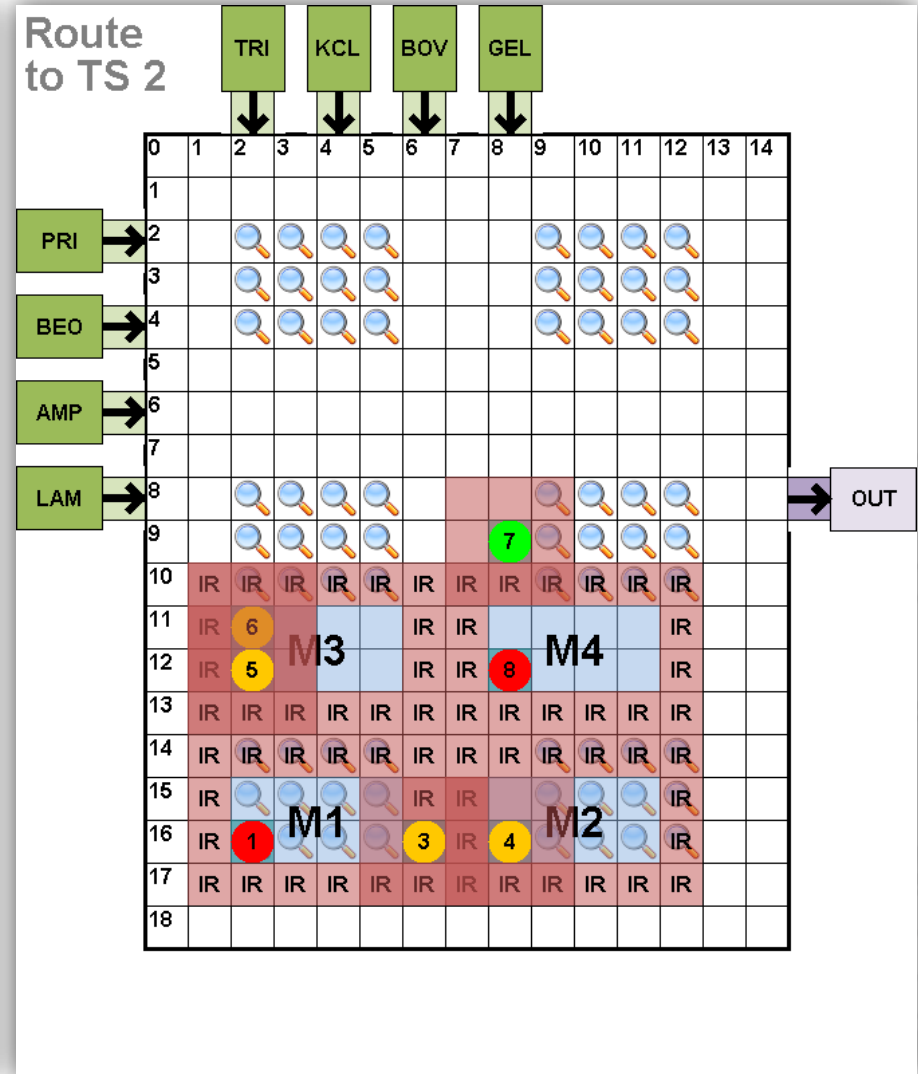


Corresponding Droplet Motion

Compaction Example

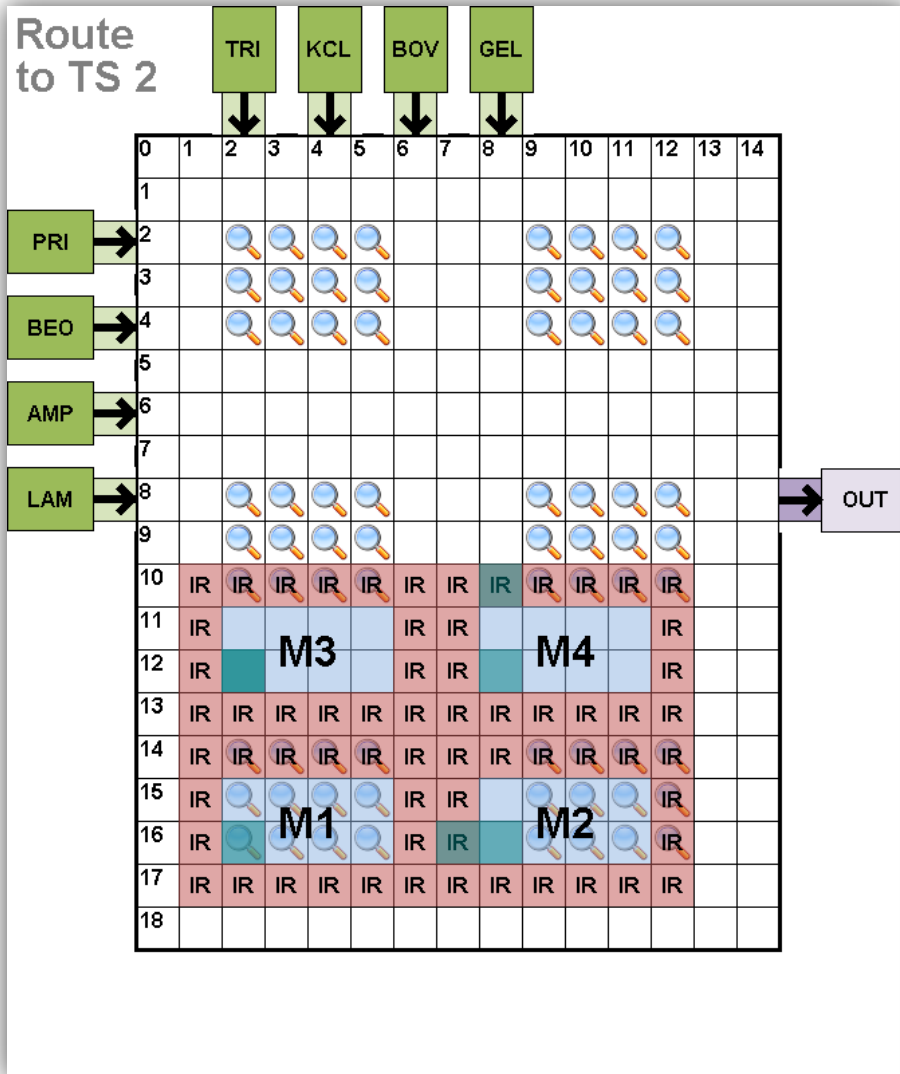


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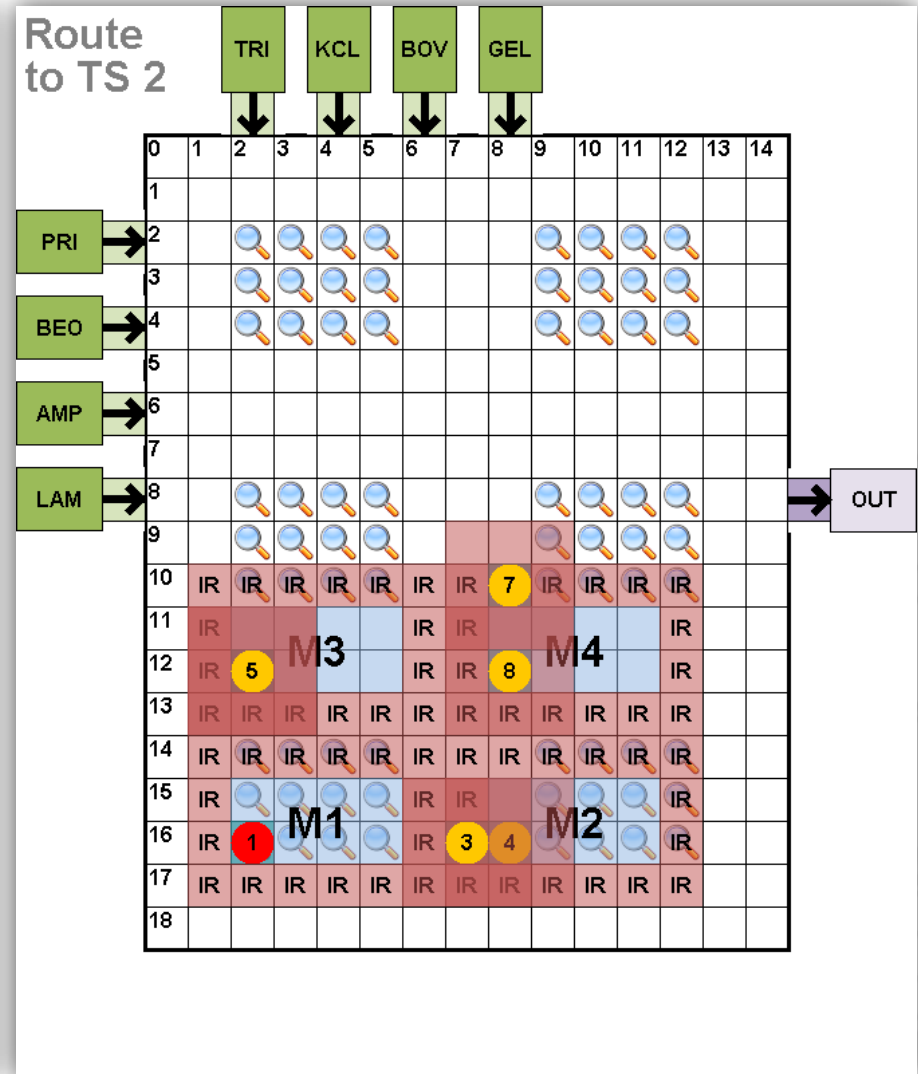


Corresponding Droplet Motion

Compaction Example

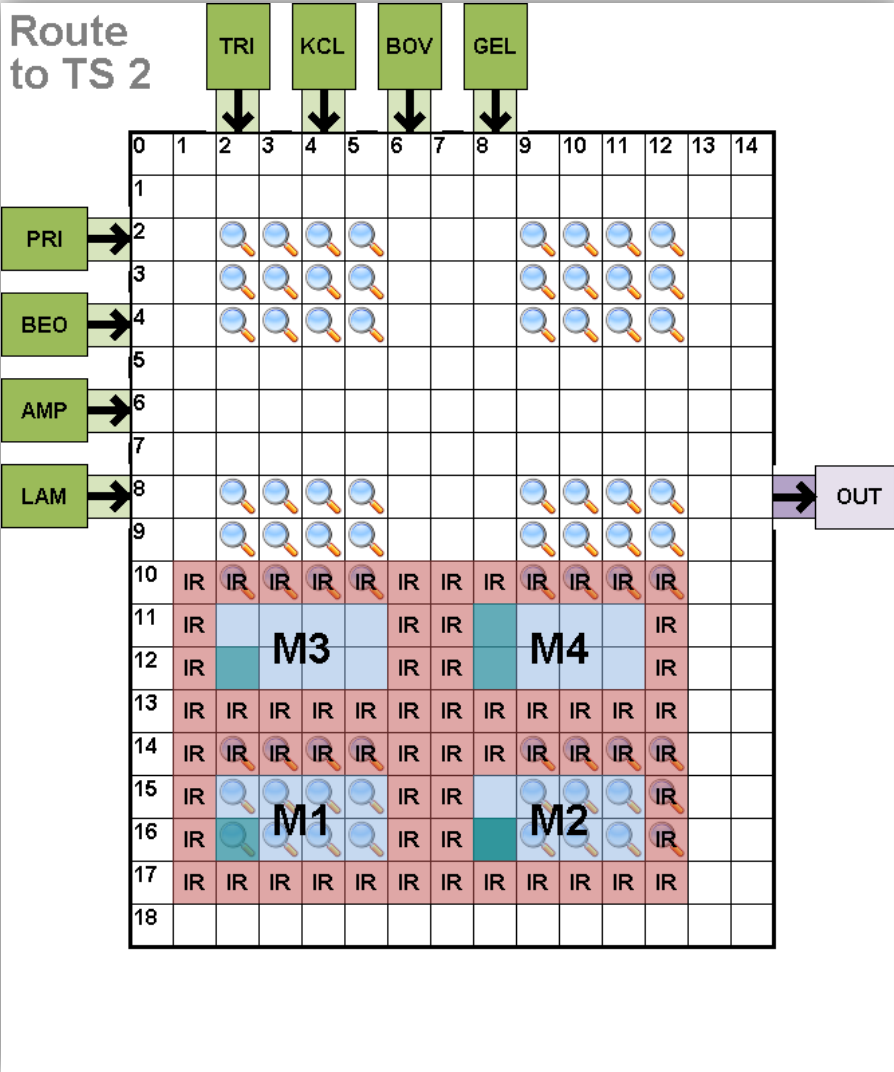


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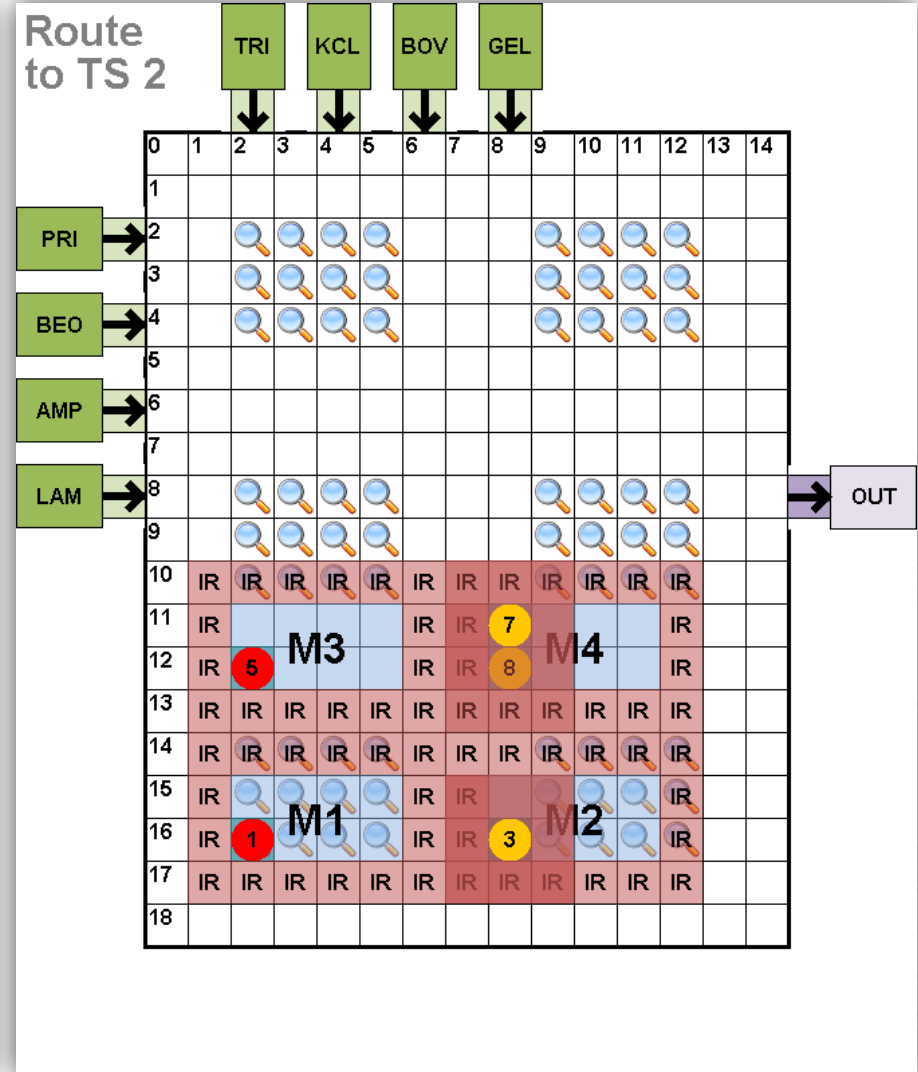


Corresponding Droplet Motion

Compaction Example

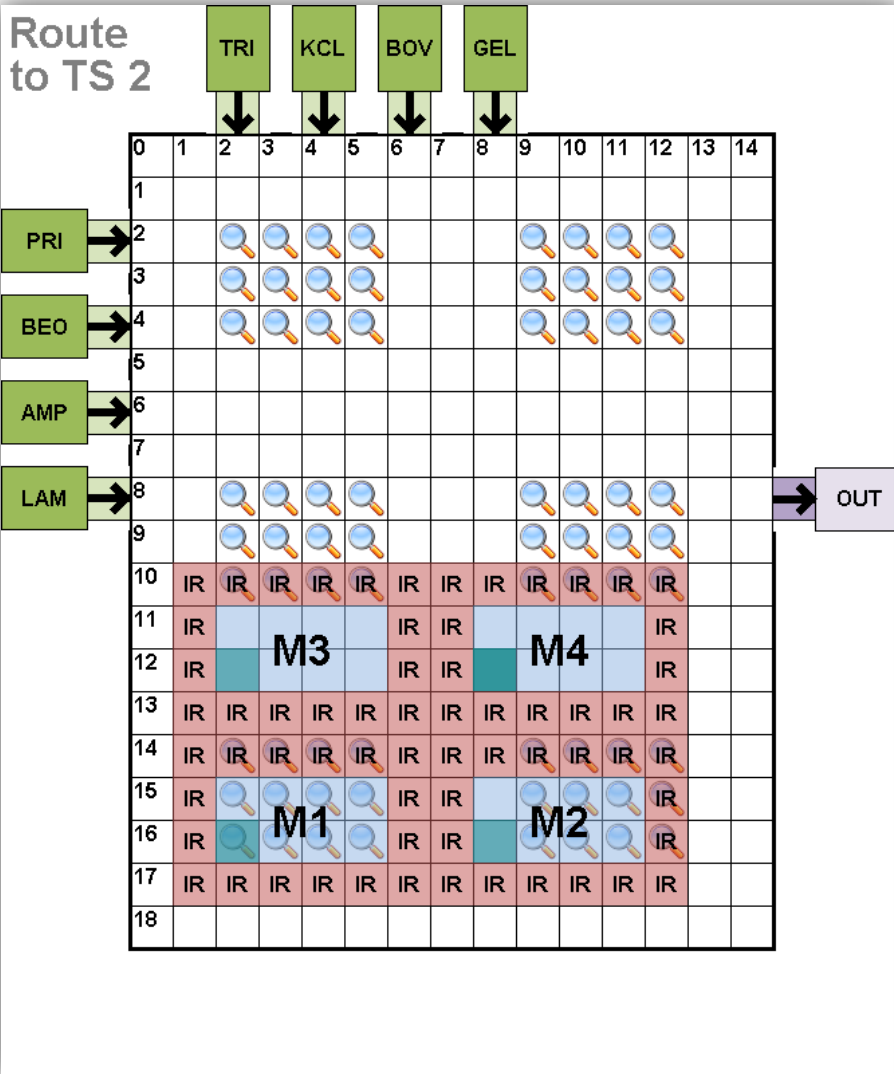


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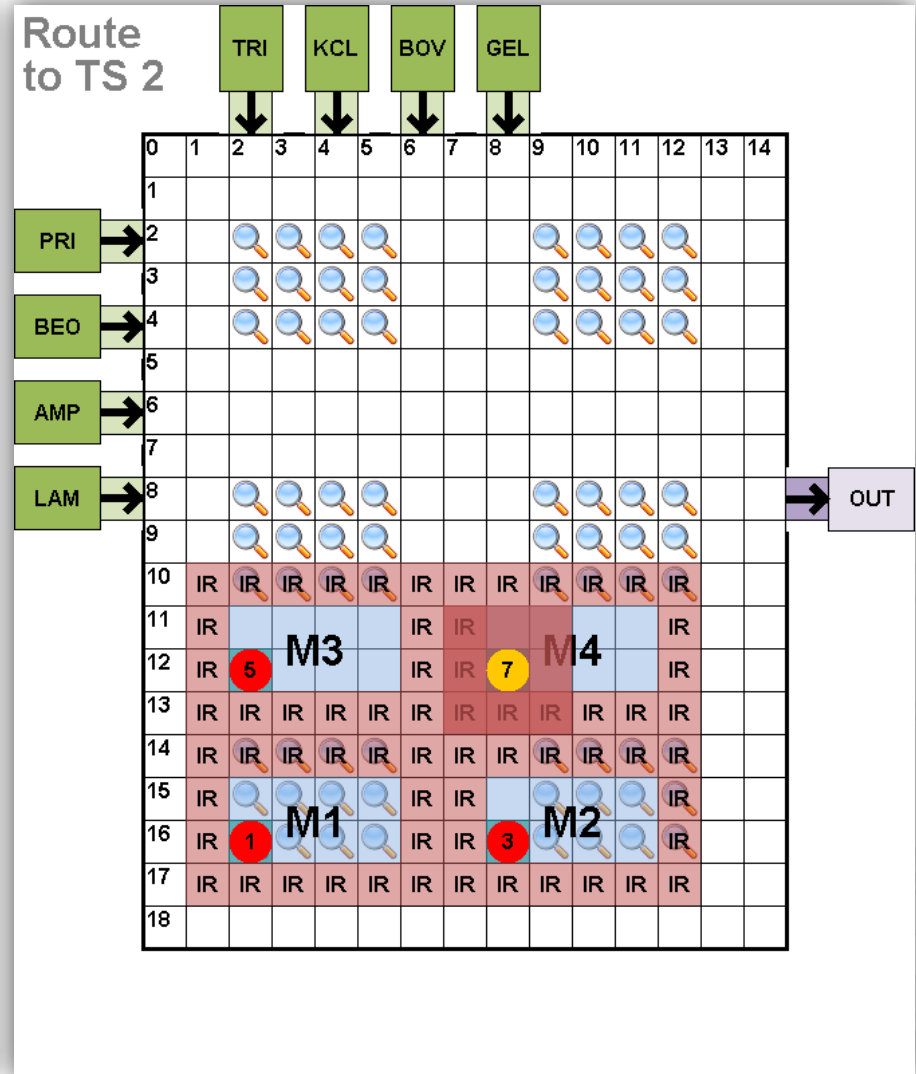


Corresponding Droplet Motion

Compaction Example

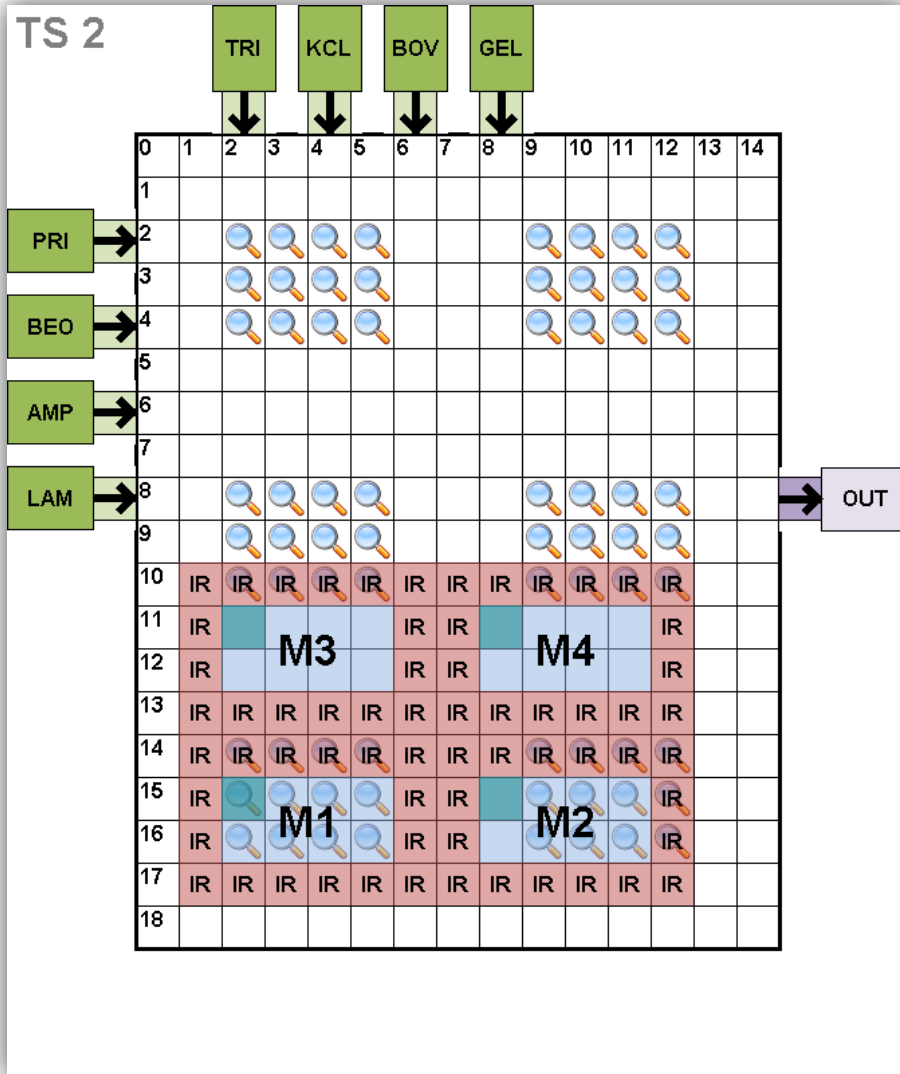


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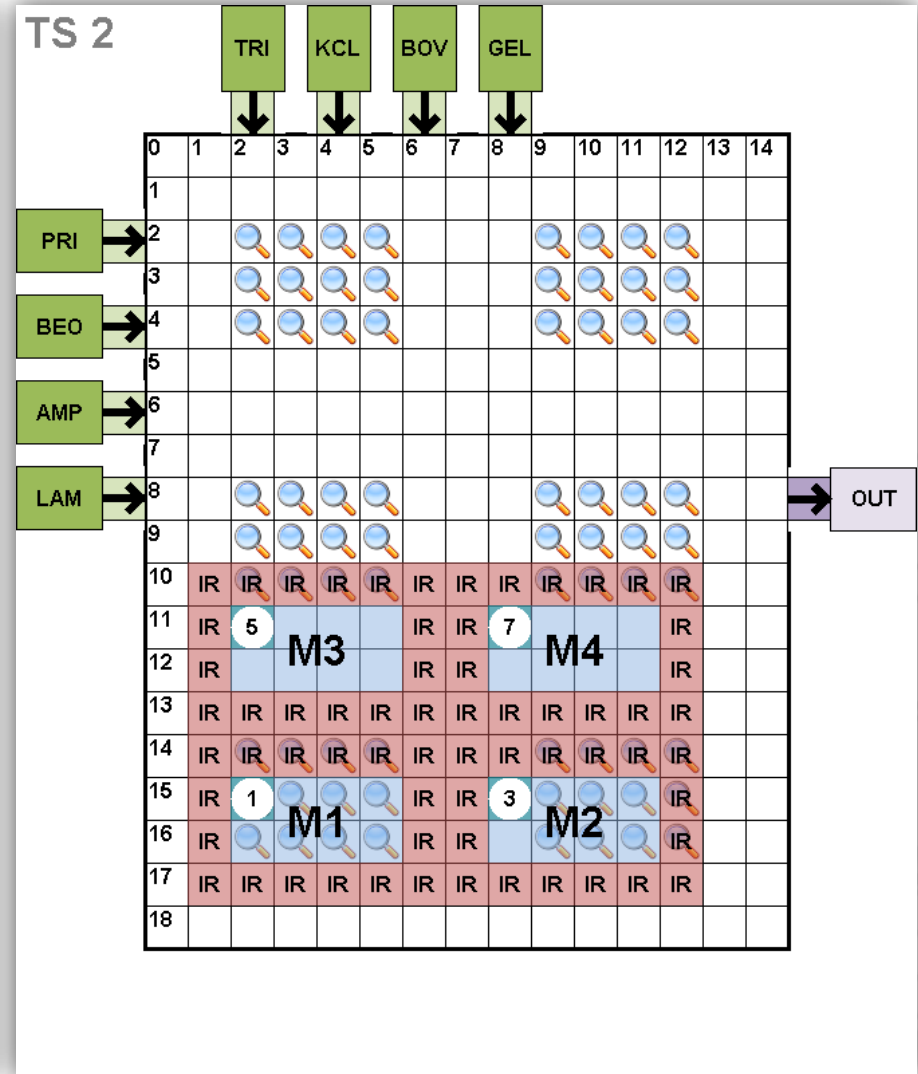


Corresponding Droplet Motion

Compaction Example

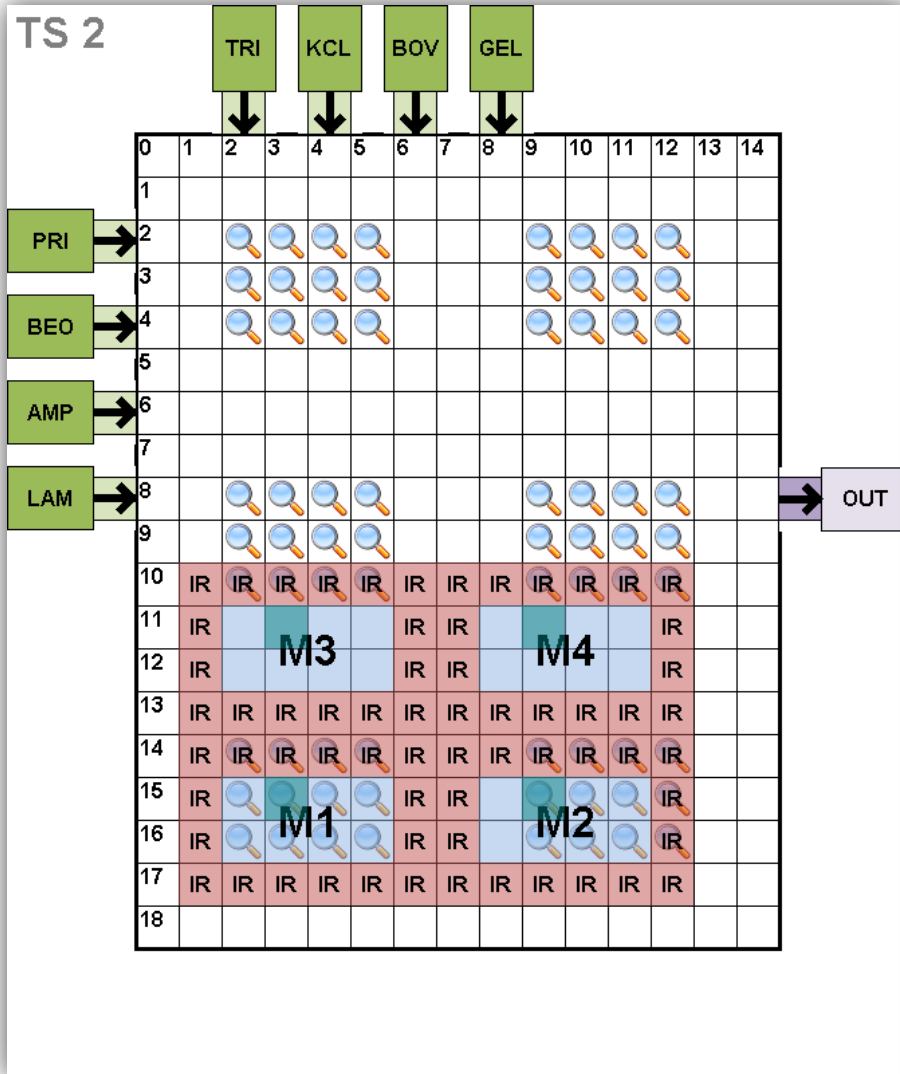


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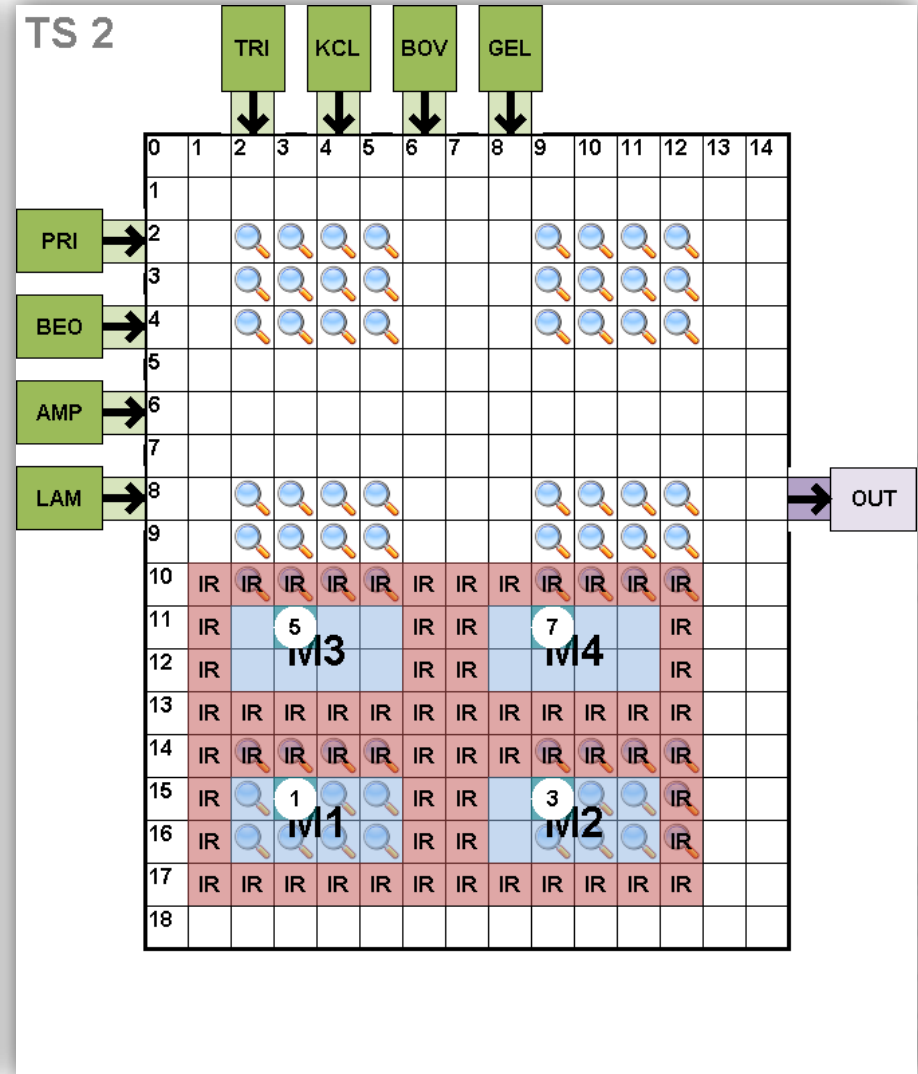


Corresponding Droplet Motion

Compaction Example

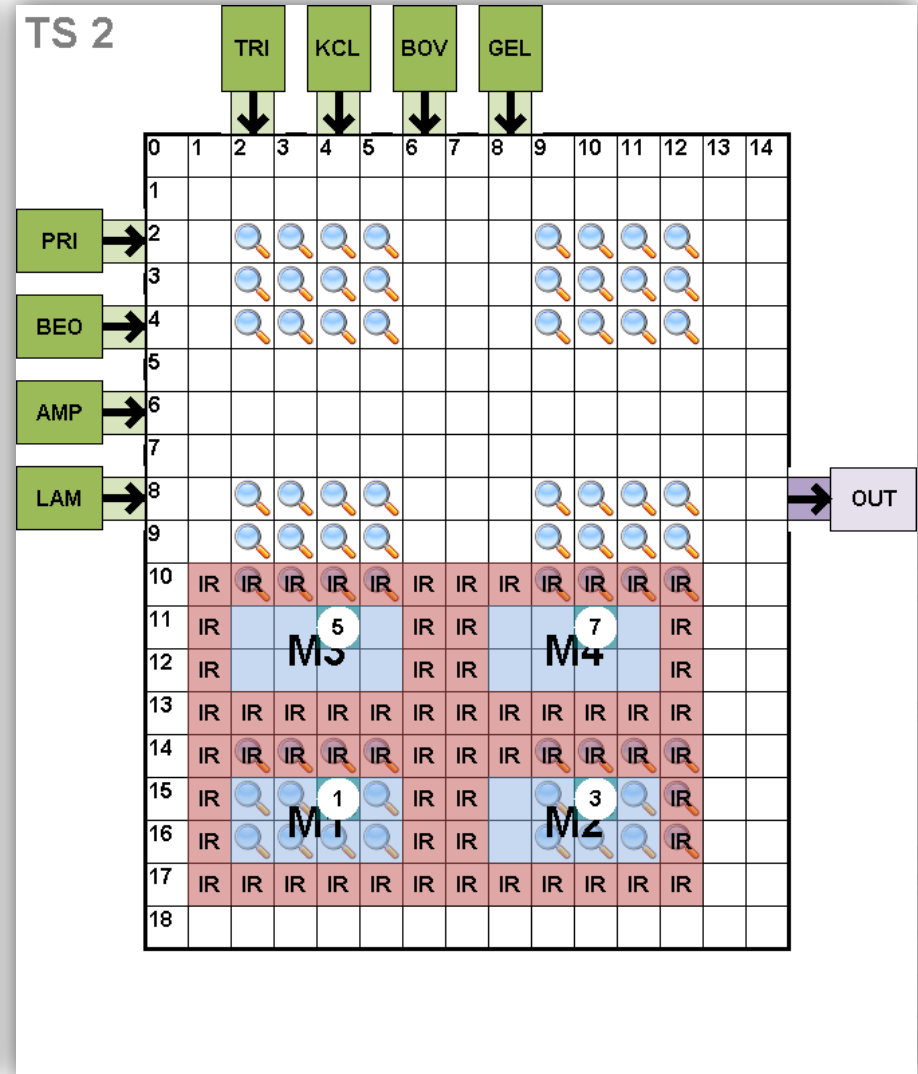
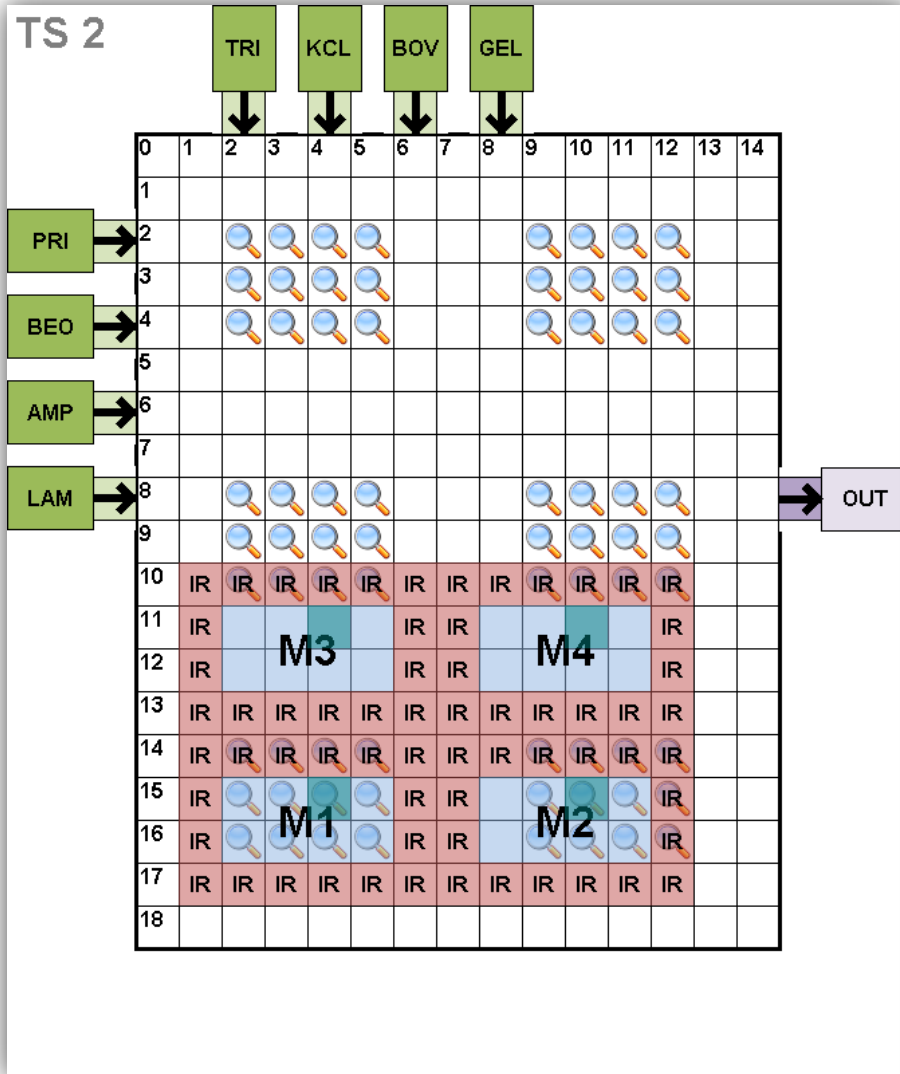


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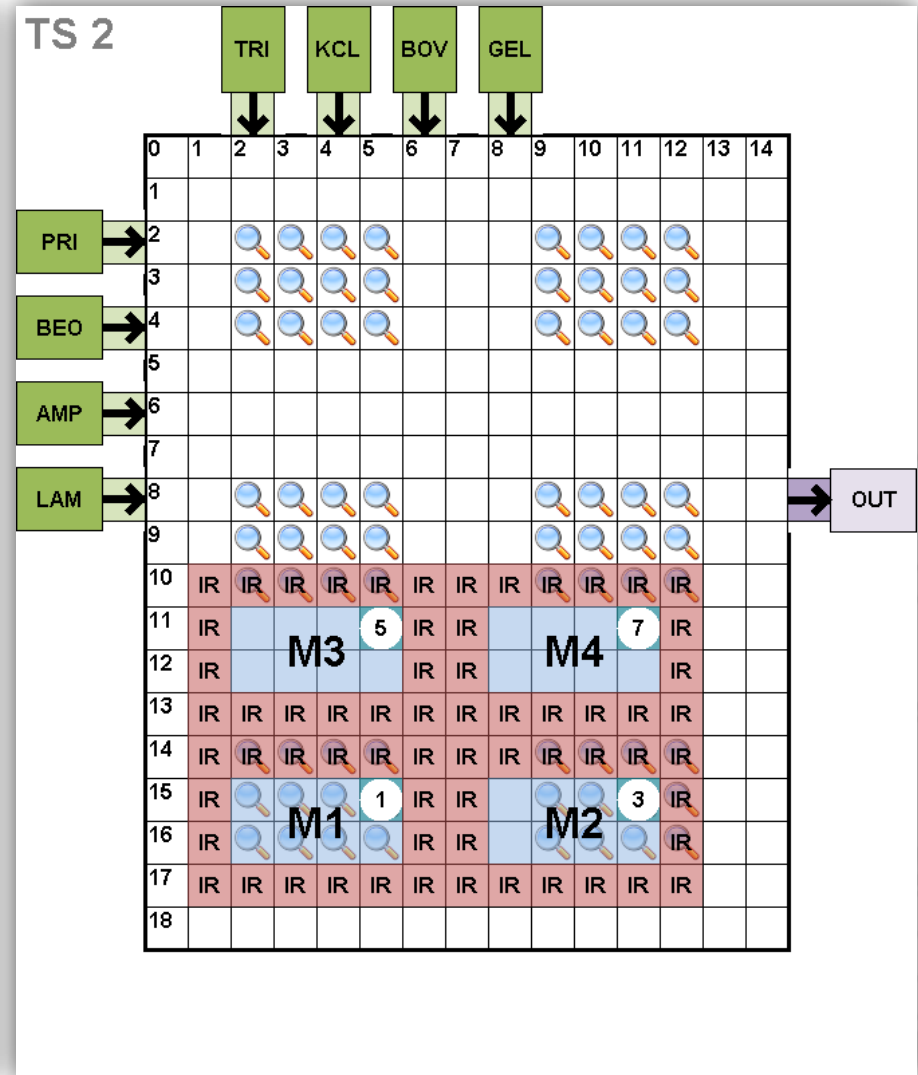
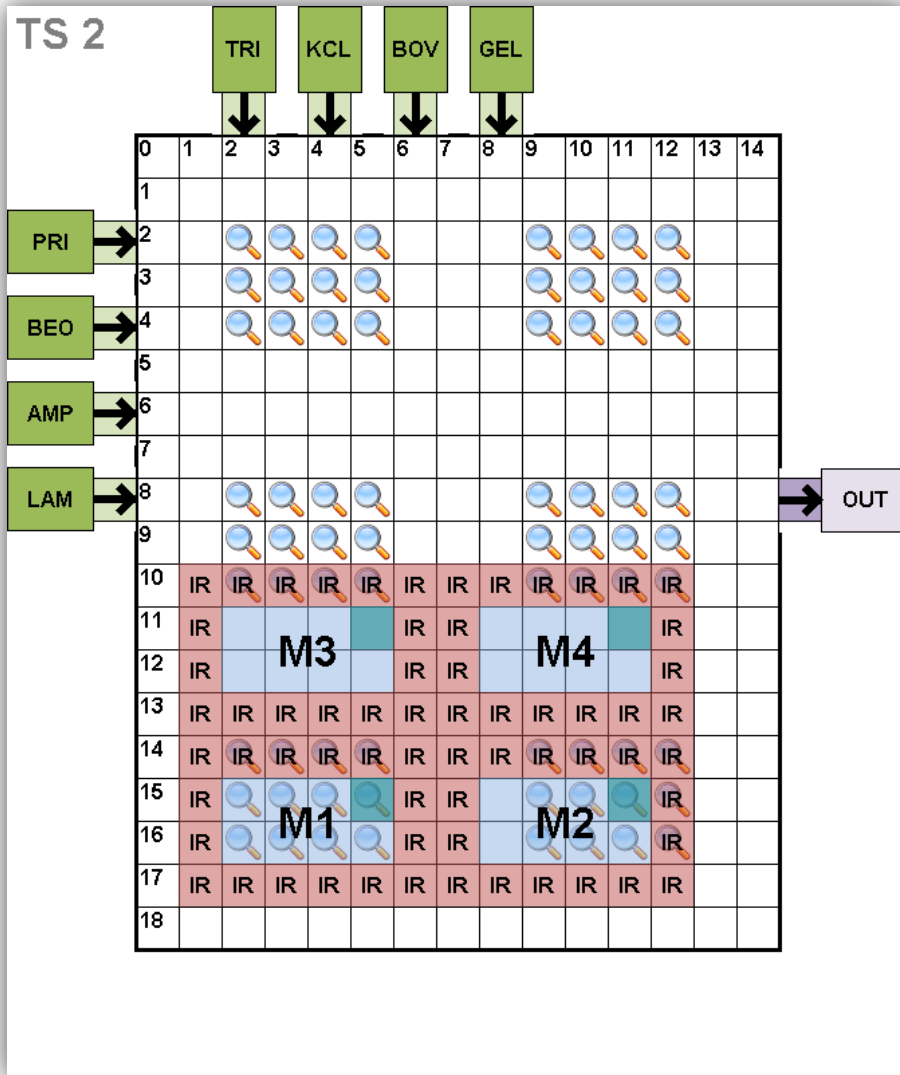


Corresponding Droplet Motion

Compaction Example

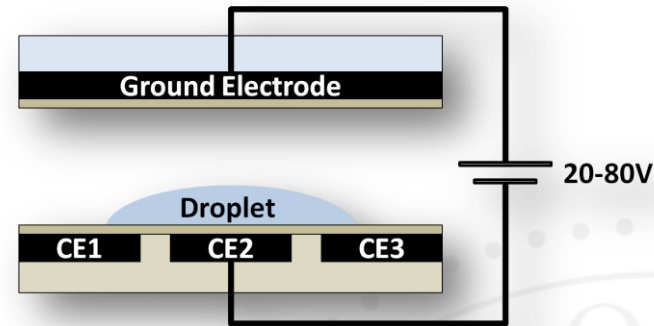
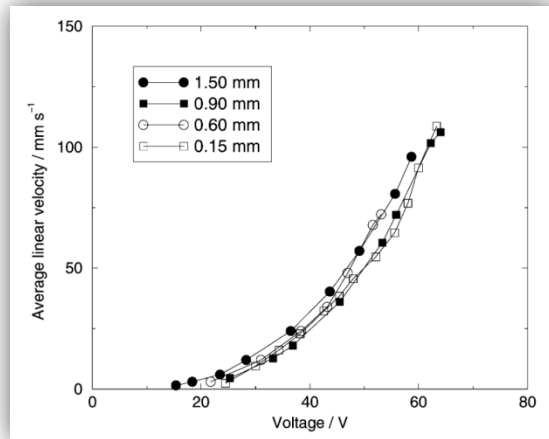


Compaction Example



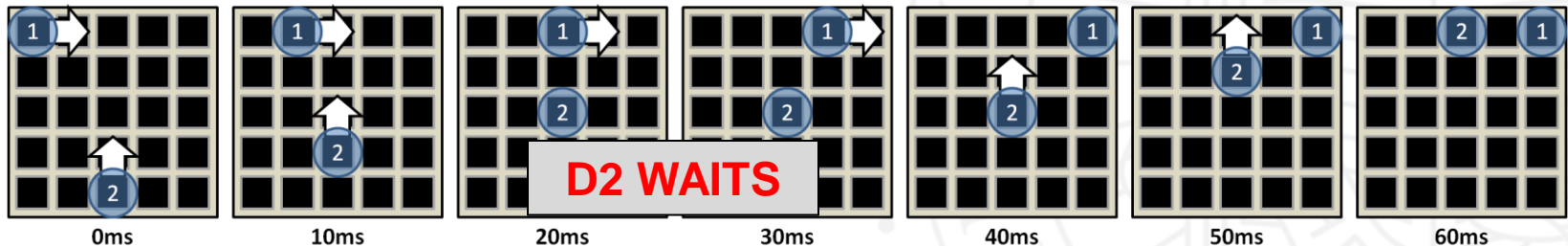
Discrete Perspective

- ▶ Increase Voltage → Increase Velocity



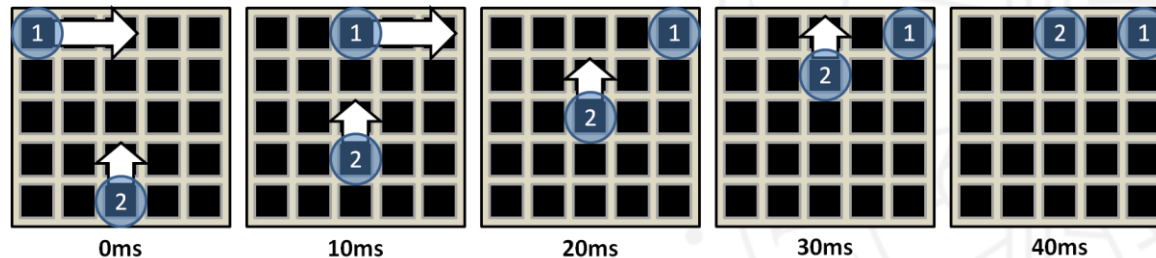
Pollack, M. G., Shenderov, A. D., and Fair, R. B. 2002. Electrowetting-based actuation of droplets for integrated microfluidics. *Lab-on-a-Chip* 2, 2 (Mar. 2002), 96-101.

- ▶ Compaction treated as discrete problem
 - ▶ Single voltage used for all droplet movements
 - ▶ All droplets move at same speed (requires halts)



Continuous-Time Perspective

- ▶ Voltages can be changed
 - ▶ Abandons synchronous droplet movement
- ▶ Reduce energy usage; maintain timing
- ▶ Compaction treated as continuous problem
 - ▶ Multiple voltages used for droplet movements
 - ▶ Droplets move at different speeds (avoid halts)



Formal Problem Formation

$$Z(d_i) = \sum_{j=1}^N z_{i,j}$$

$$T(d_i) = \sum_{j=1}^N (t_{i,j} + u_{i,(j,j+1)})$$

$$U(d_i, p_{i,j}) = \sum_{k=0}^{j-1} (t_{i,k} + u_{i,(k,k+1)})$$

$$I(d_i, p_{i,j}) = [U(d_i, p_{i,j}), U(d_i, p_{i,j}) + t_{i,j}]$$

$$J(d_i, p_{i,j}, p_{i,j+1}) = (U(d_i, p_{i,j}) + t_{i,j}, U(d_i, p_{i,j+1}))$$

$$Q(d_i) = \langle I(d_i, p_{i,1}), J(d_i, p_{i,1}, p_{i,2}), \dots, I(d_i, p_{i,N-1}, p_{i,N}) \rangle$$

$$C = (K - 1) \sum_{i=1}^K C_i$$

$$\text{Velocity} = 0.995 \times \text{Voltage}^2$$

$$+ 0.0358 \times \text{Voltage} - 0.9103$$

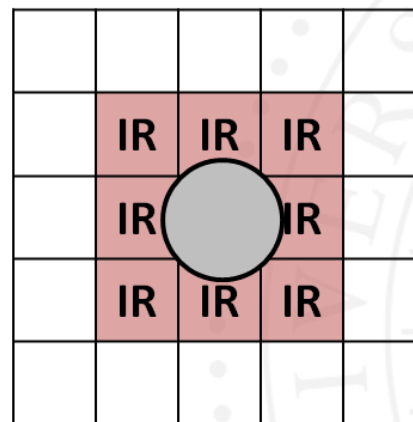
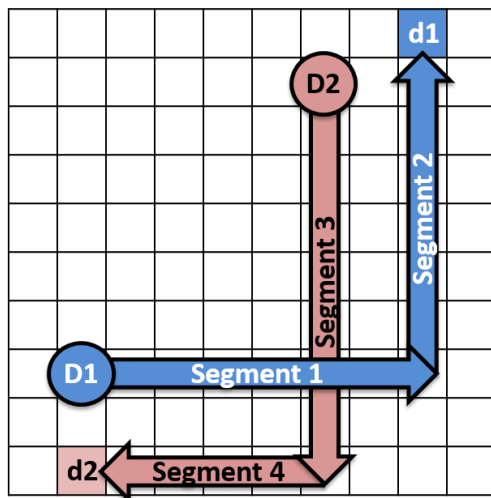
$$\text{Energy} = \text{Power} \times \text{Time} = \frac{\text{Voltage}^2}{\text{Resistance}} \times \text{Time}$$

$$E_{i,j} = \frac{V_{i,j}^2}{1G\Omega} \times \Delta T_{i,j}$$

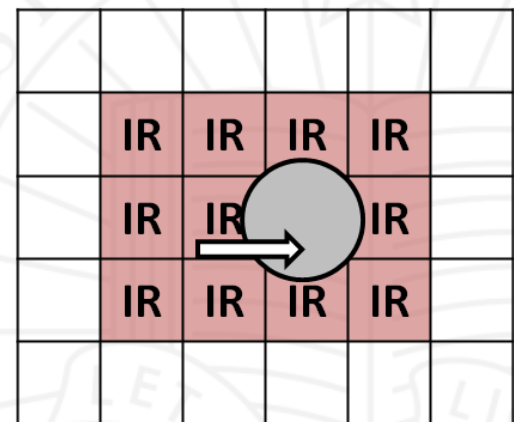
Details In Paper

General Problem Formation

- Droplet paths broken into segments
 - Max-length contiguous subsequence in one direction
- Droplet motion:
 - Constant velocity/voltage along entire segment
 - Only stops at beginning/end of segments
 - Interference constraints at continuous-time positions



Static Constraints

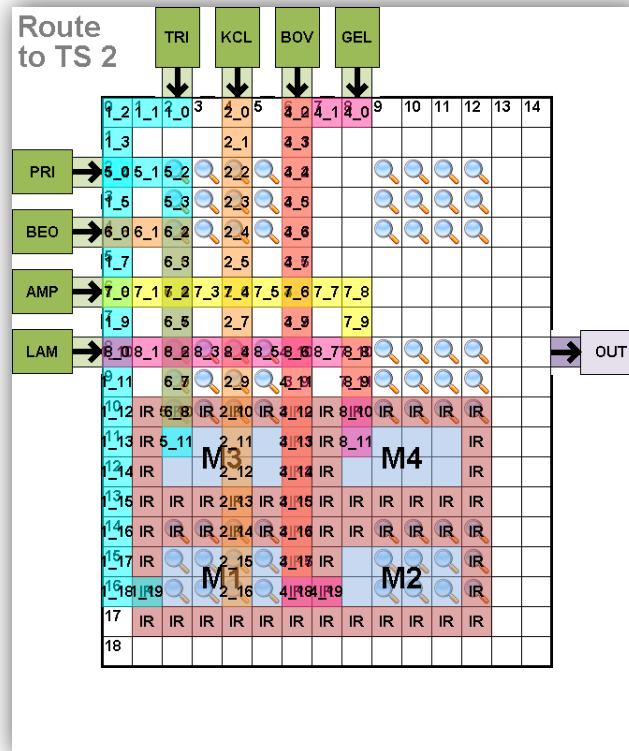


Dynamic Constraints

Interference Regions (IR) Prevent Droplet Collisions

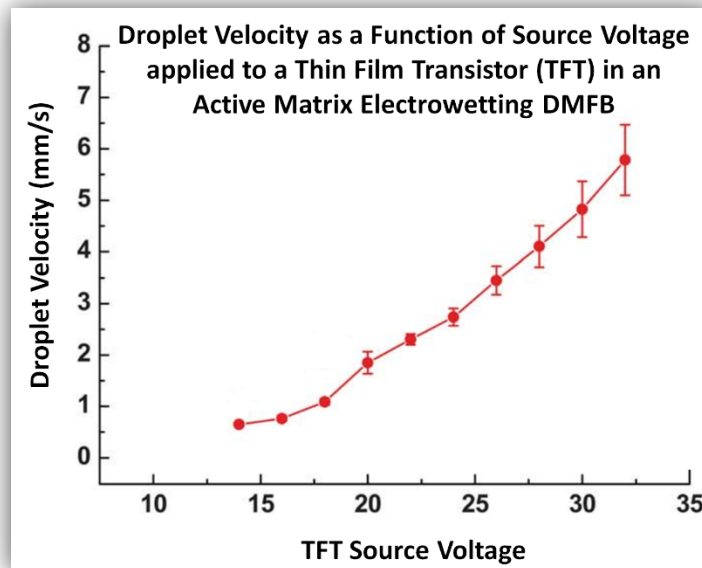
Algorithmic Description

- ▶ Step 1: Route computation
 - ▶ Roy's maze-based droplet router (greedy)
 - ▶ Computes routes that could overlap
 - ▶ Never re-visit/re-compute routes



Algorithmic Description

- Step 2: Time-constrained, energy-aware compaction
 - Given timing constraint T_c
 - For each droplet path:
 - Compute initial path velocity $vel = \frac{pathLength}{T_c}$
 - Minimum Voltage for velocity derived from graph



$$Velocity = 0.005 \times Voltage^2 + 0.0358 \times Voltage - 0.9103$$

Least-squares-fit equation

Noh, J. H., Noh, J., Kreit, E., Heikenfeld, J., and Rack, P. D. 2012. Toward active-matrix lab-on-a-chip: programmable electrofluidic control enabled by arrayed oxide thin film transistors. *Lab-on-a-Chip* 12, 2 (Jan. 2012), 353-360.

Algorithmic Description

Step 2: Compaction (continued)

Compute all segment timings from initial velocities

For each droplet path P_d

For each electrode position e_{di} in P_d

Compare against each previously compacted path

If no interference along segment:

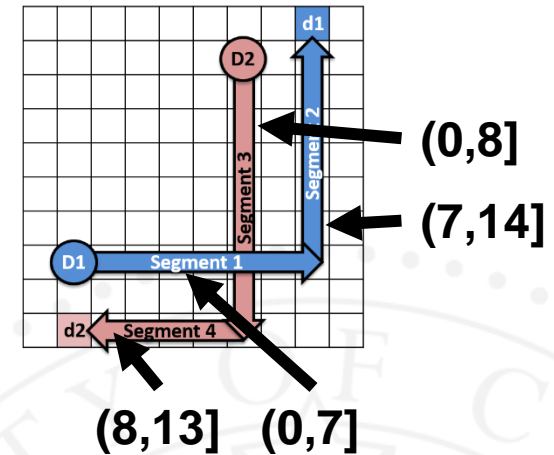
Accept segment

If interference along segment:

Speedup current droplet along its segment

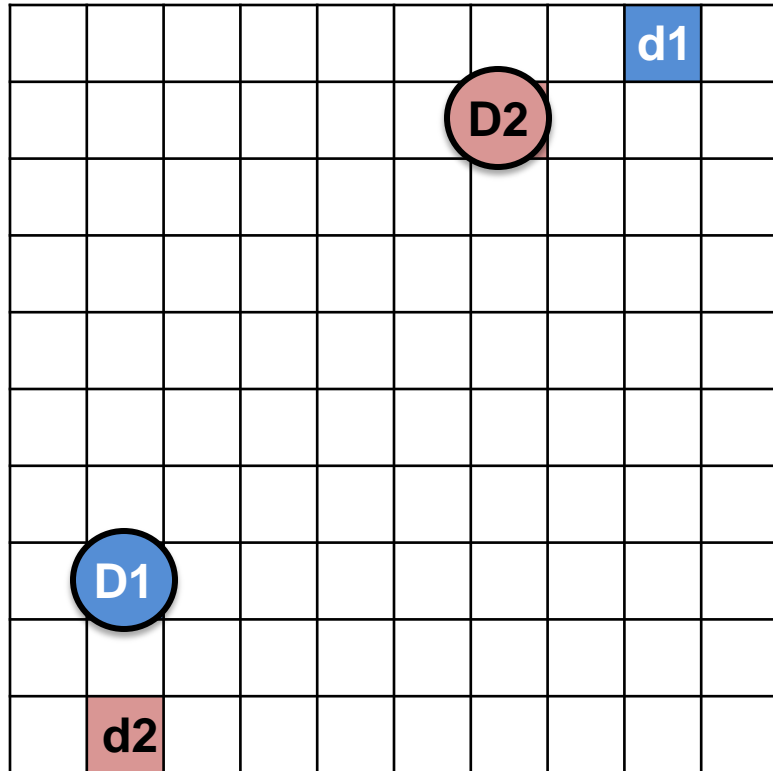
Adjust remaining segments to conserve energy

Re-compute path timings for that droplet

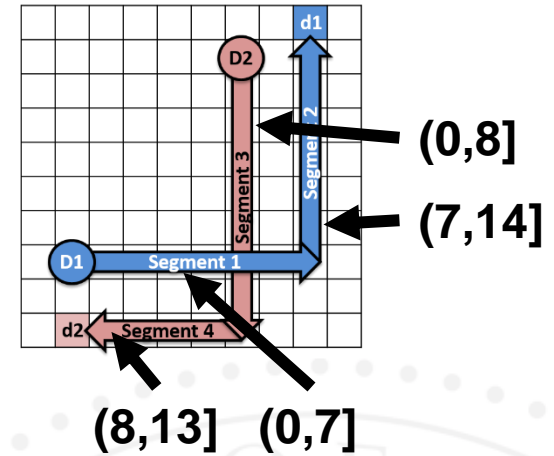


Example Coming!

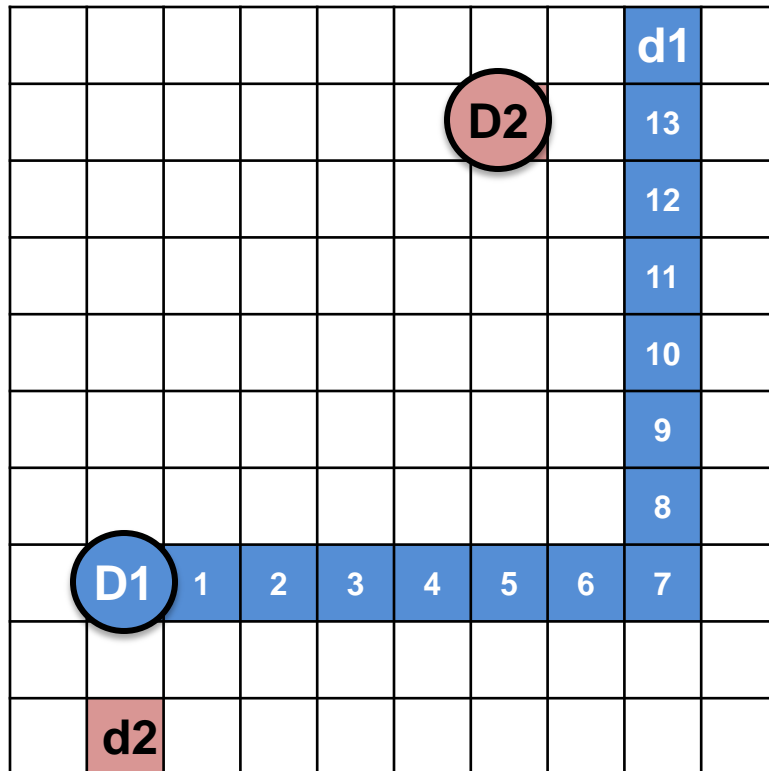
Simple Example



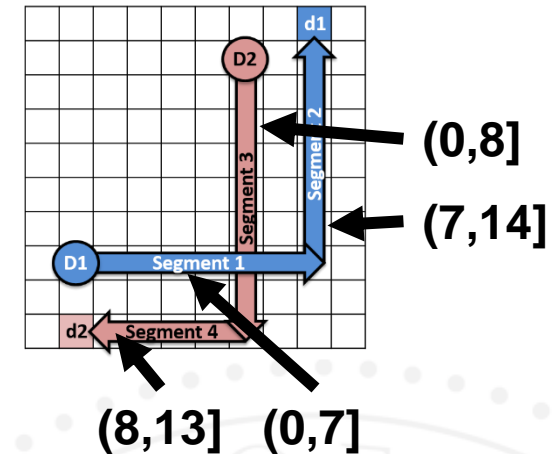
Compact D1.



Simple Example



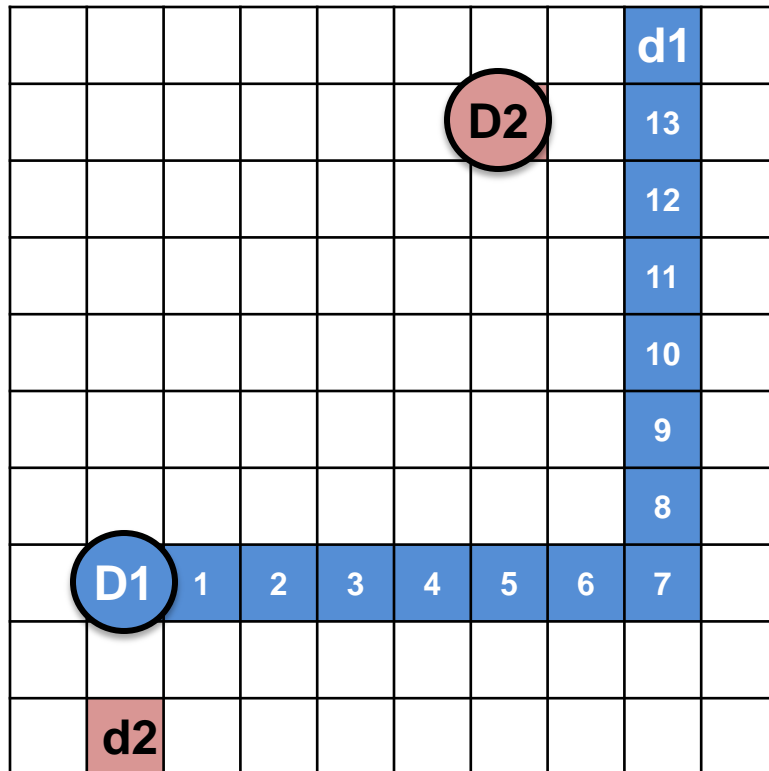
No previous paths; D1 routes with no problems.



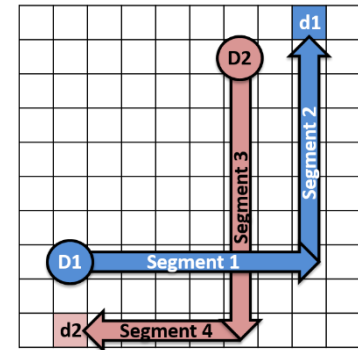
Numbers on electrodes indicate the time the droplet arrives at the electrode.

Segment 1: 1 electrode/s
Segment 2: 1 electrode/s

Simple Example



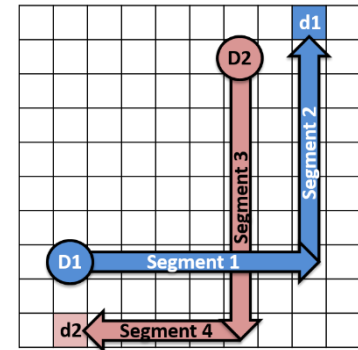
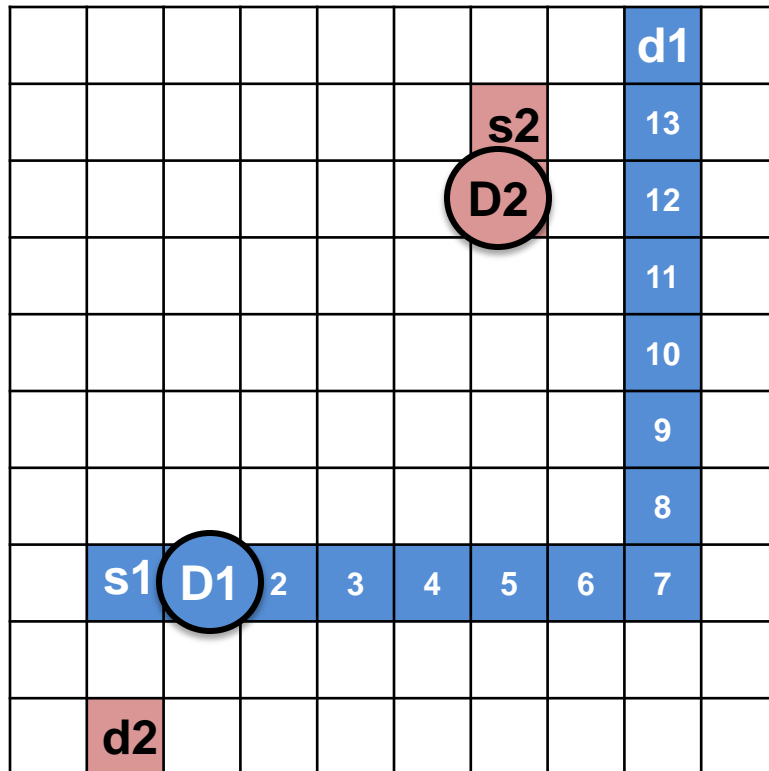
Now compact D2 against all previous droplet paths (D1).



Numbers on electrodes indicate the time the droplet arrives at the electrode.

Segment 1: 1 electrode/s
Segment 2: 1 electrode/s
Segment 3: 1 electrodes/s

Simple Example

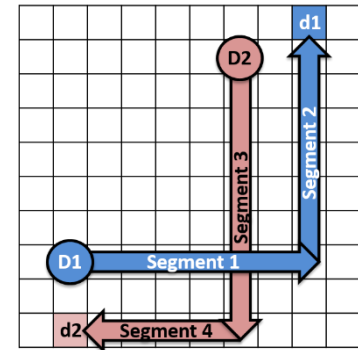
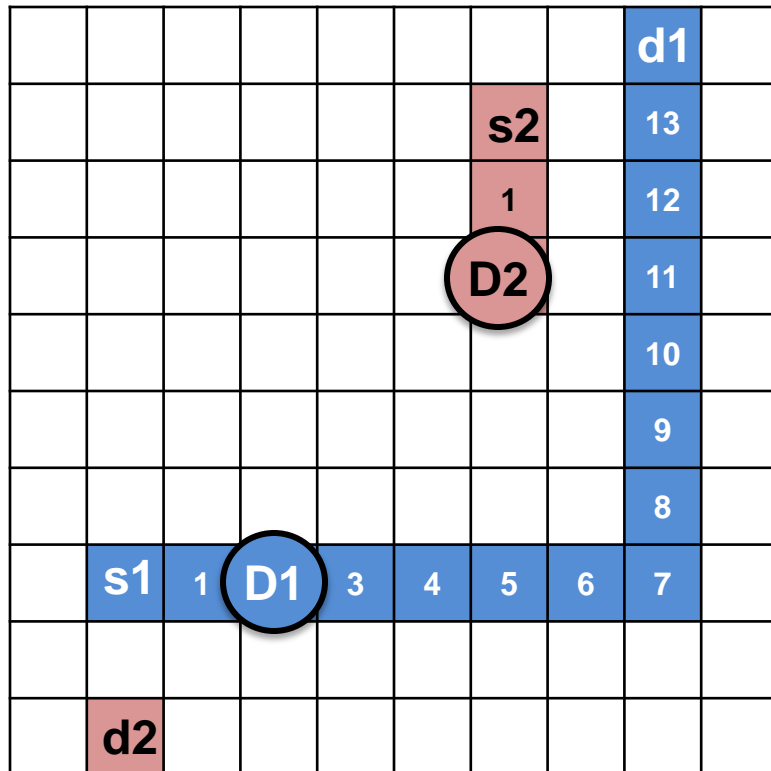


Numbers on electrodes indicate the time the droplet arrives at the electrode.

Segment 1: 1 electrode/s
 Segment 2: 1 electrode/s
 Segment 3: 1 electrodes/s

Now compact D2 against all previous droplet paths (D1).

Simple Example

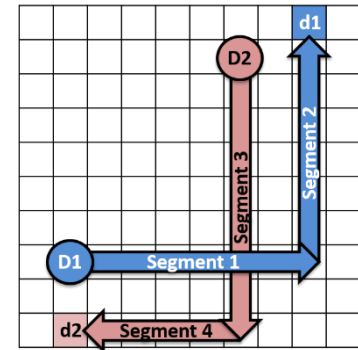
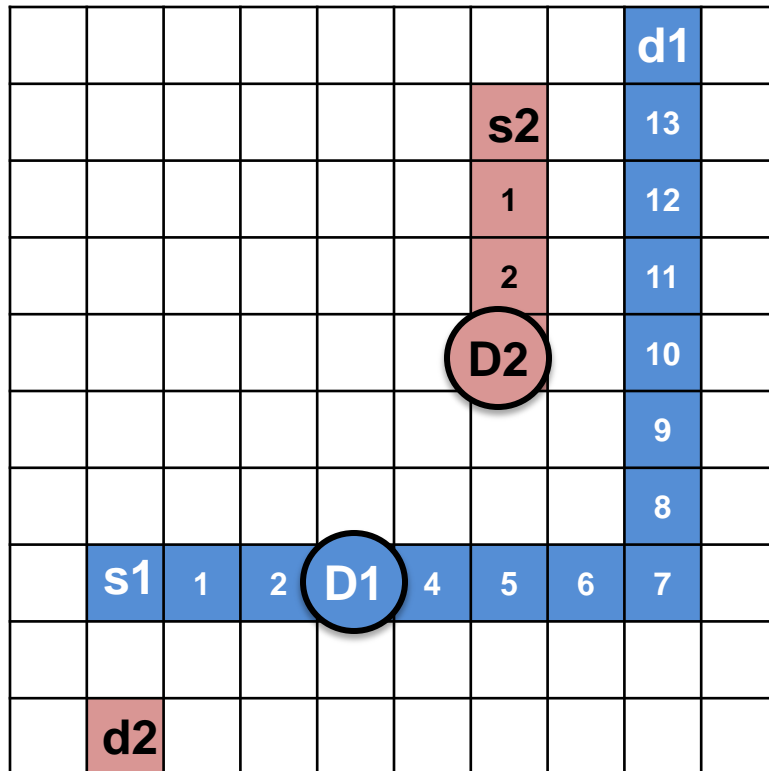


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Segment 1: 1 electrode/s
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 Segment 3: 1 electrodes/s

Now compact D2 against all previous droplet paths (D1).

Simple Example

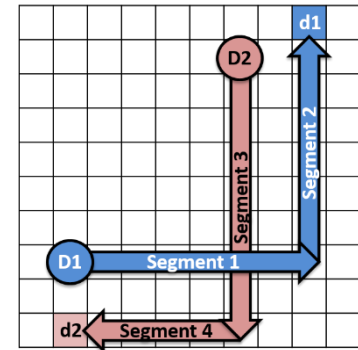
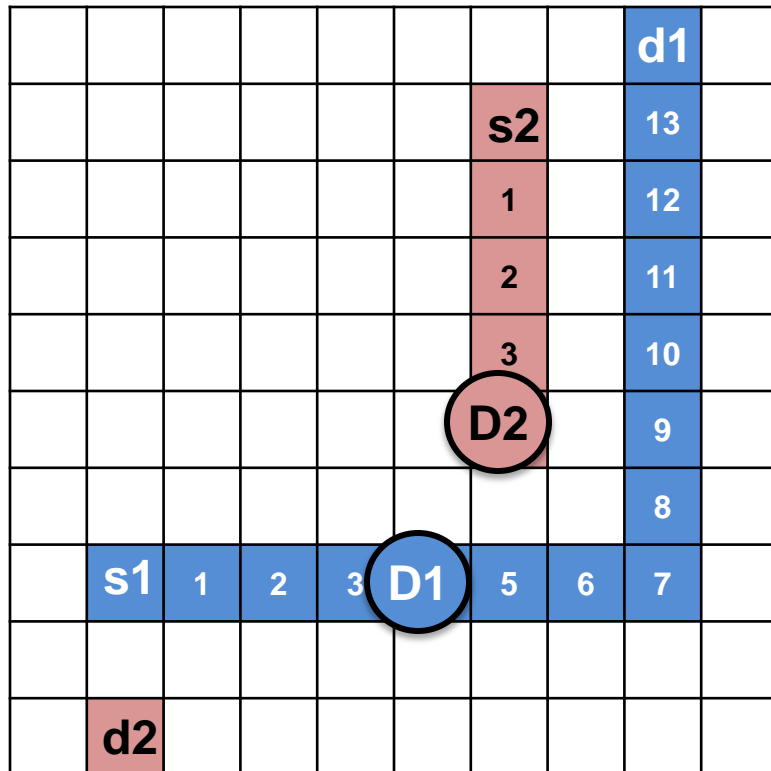


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 Segment 3: 1 electrodes/s

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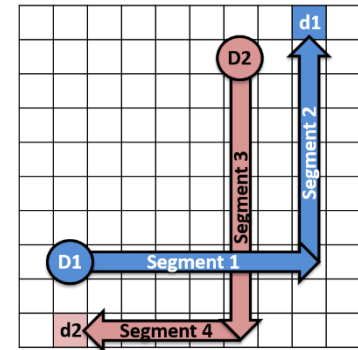
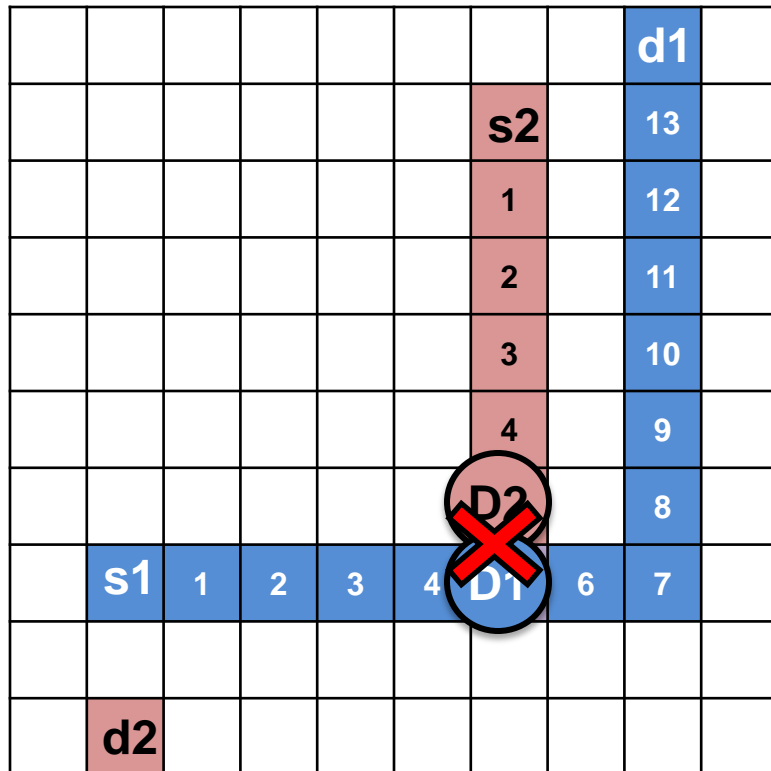


Numbers on electrodes indicate the time the droplet arrives at the electrode.

Segment 1: 1 electrode/s
 Segment 2: 1 electrode/s
 Segment 3: 1 electrodes/s

Now compact D2 against all previous droplet paths (D1).

Simple Example

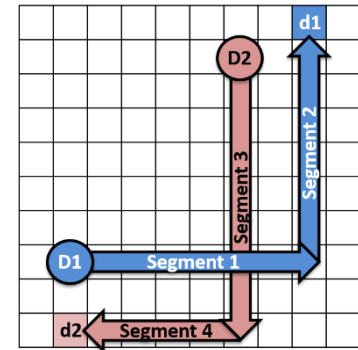
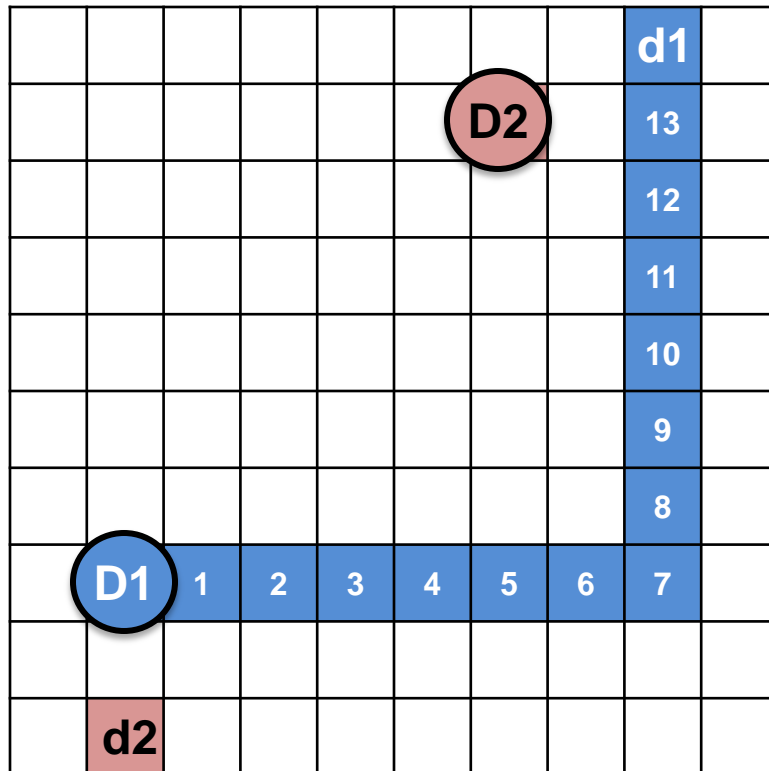


Numbers on electrodes indicate the time the droplet arrives at the electrode.

Segment 1: 1 electrode/s
 Segment 2: 1 electrode/s
 Segment 3: 1 electrodes/s

While compacting D2, detected interference at time 5 between D1 and D2.

Simple Example

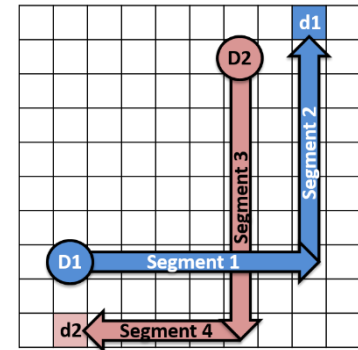
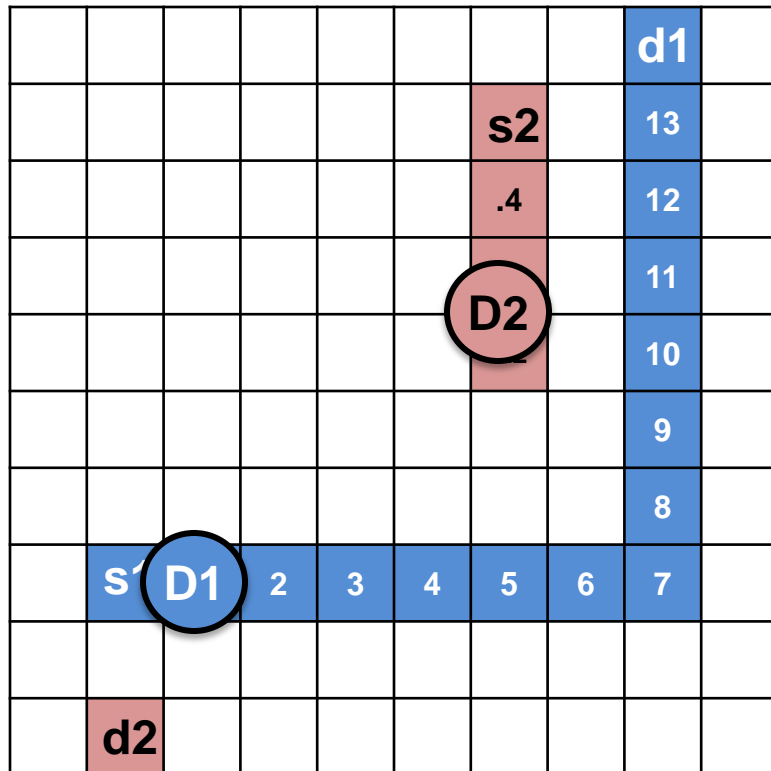


Numbers on electrodes indicate the time the droplet arrives at the electrode.

Segment 1: 1 electrode/s
Segment 2: 1 electrode/s
Segment 3: 2.5 electrodes/s

Increases D2's velocity/voltage (2.5x) and restart compaction for D2.

Simple Example

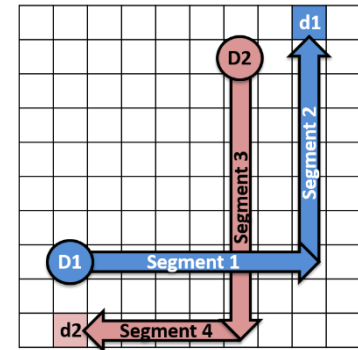
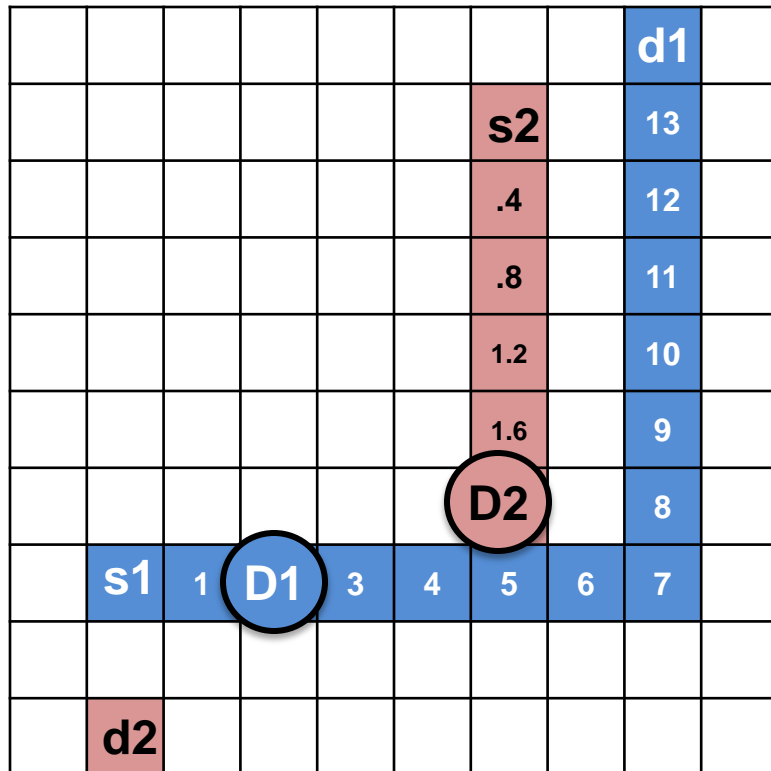


Numbers on electrodes indicate the time the droplet arrives at the electrode.

Segment 1: 1 electrode/s
 Segment 2: 1 electrode/s
 Segment 3: 2.5 electrodes/s

Re-compact D2 at 2.5x speed against all previous droplet paths (D1).

Simple Example

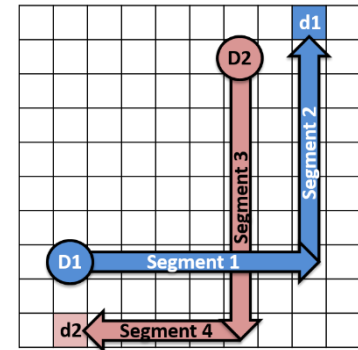
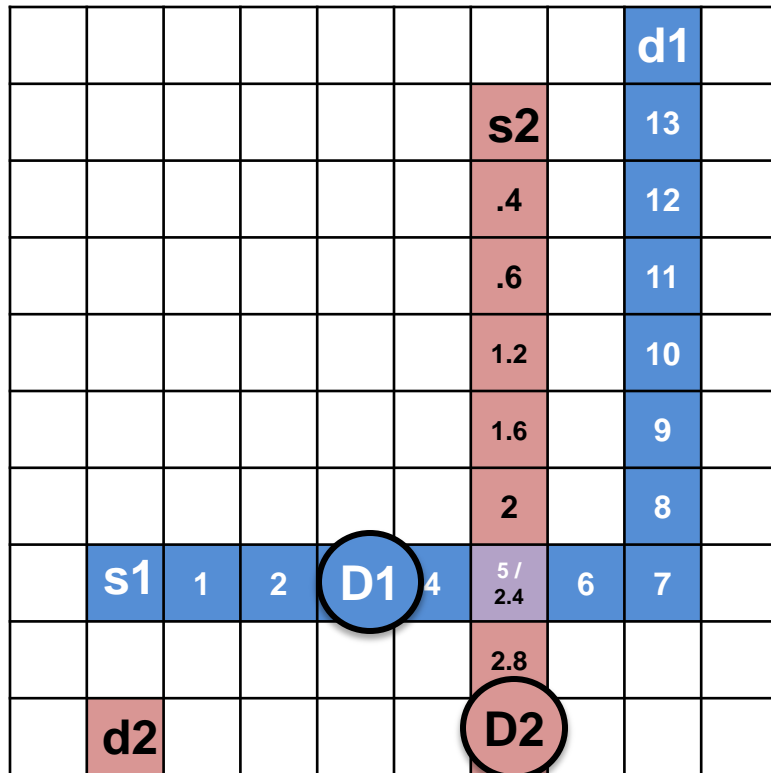


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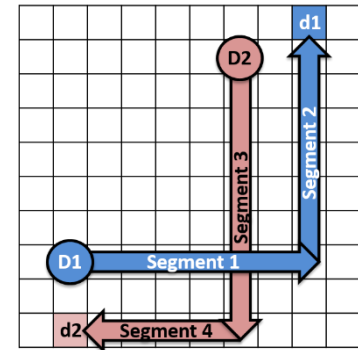
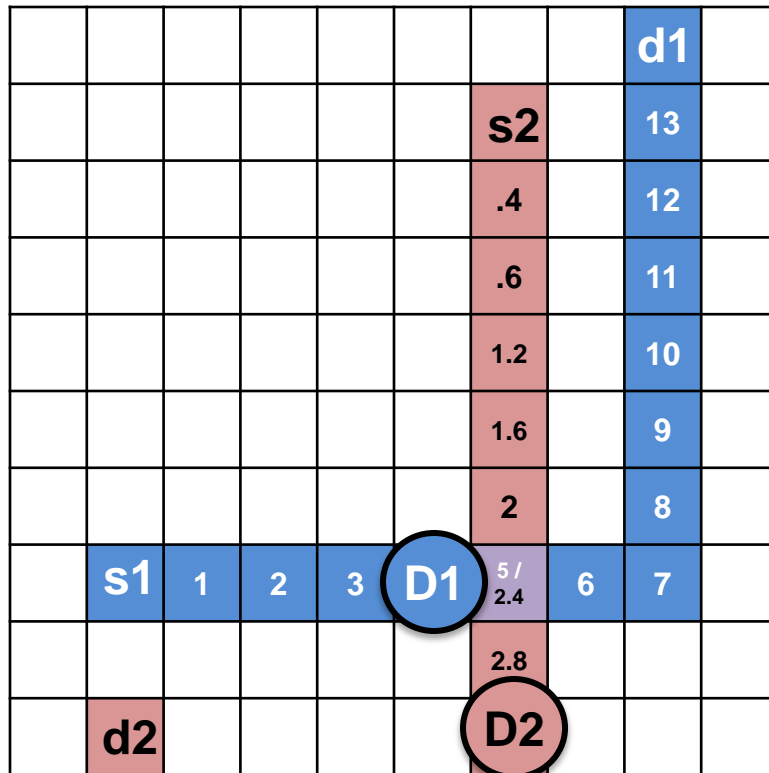


Numbers on electrodes indicate the time the droplet arrives at the electrode.

Segment 1: 1 electrode/s
 Segment 2: 1 electrode/s
 Segment 3: 2.5 electrodes/s

D2 reached end of segment.

Simple Example

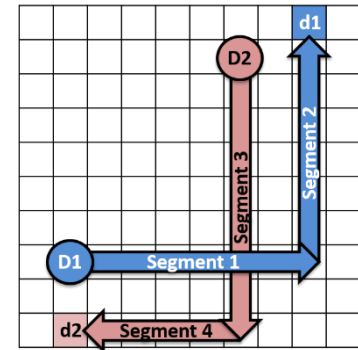
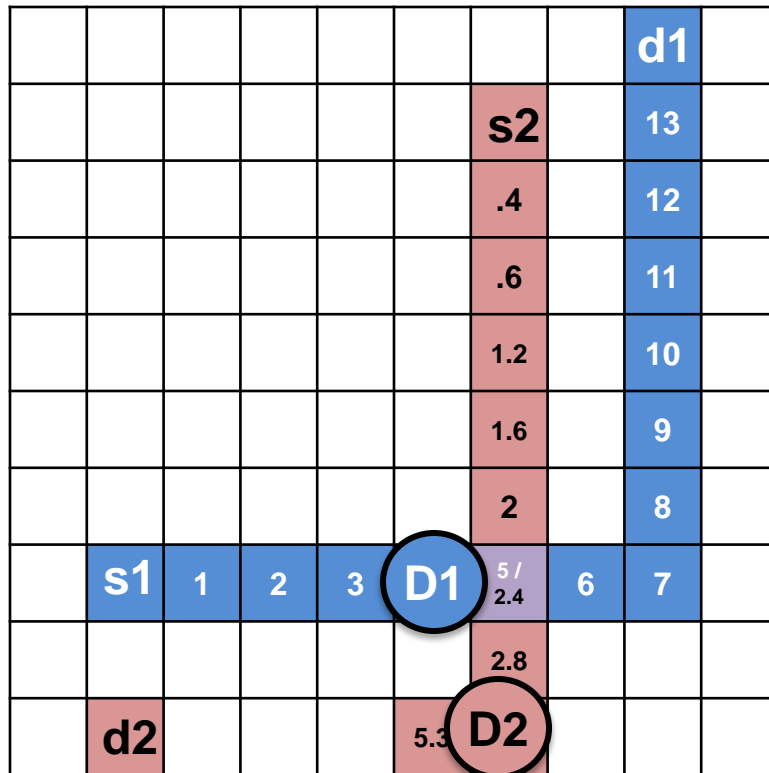


Numbers on electrodes indicate the time the droplet arrives at the electrode.

Segment 1: 1 electrode/s
 Segment 2: 1 electrode/s
 Segment 3: 2.5 electrodes/s
 Segment 4: 0.46 electrodes/s

D2 does not need to get there before D1; save energy and slow D2 down to 0.46 electrodes/sec.

Simple Example

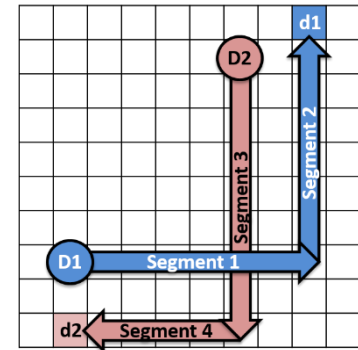
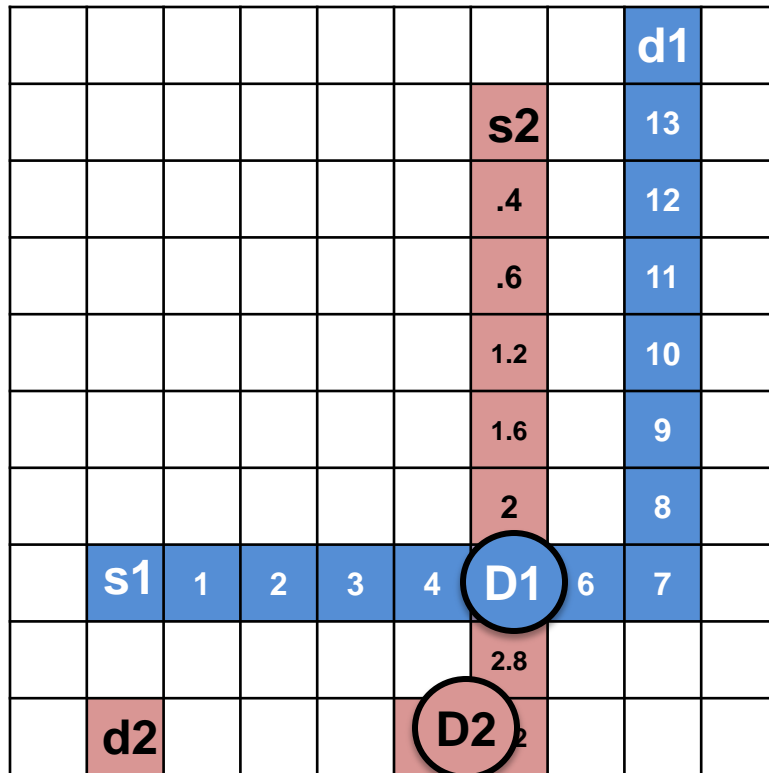


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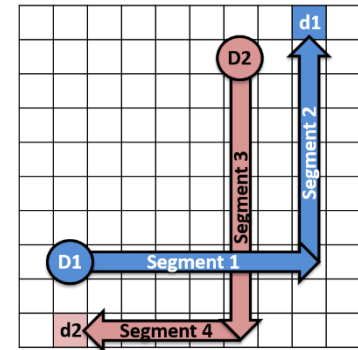
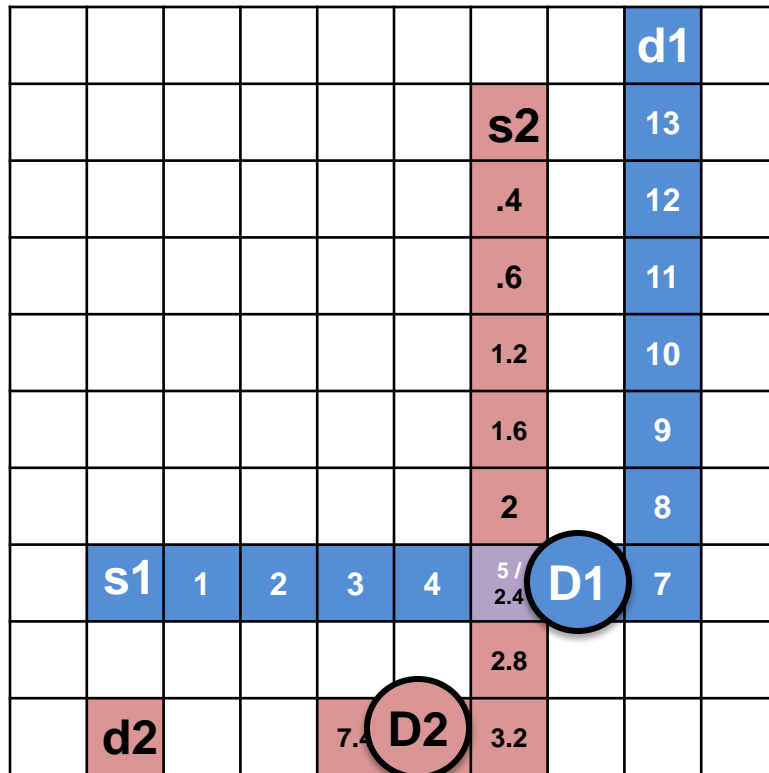


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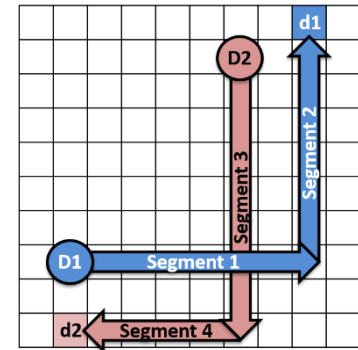
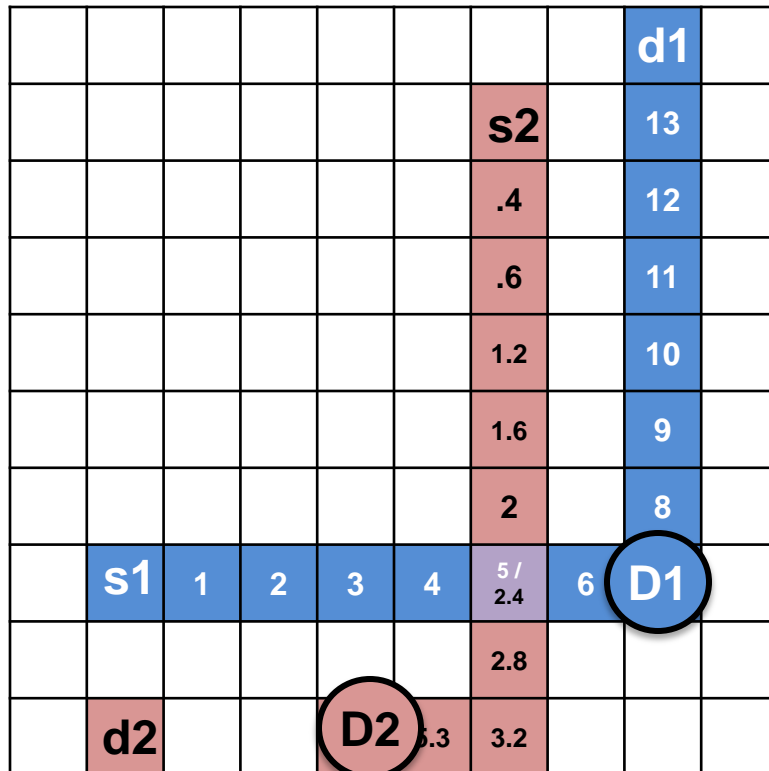


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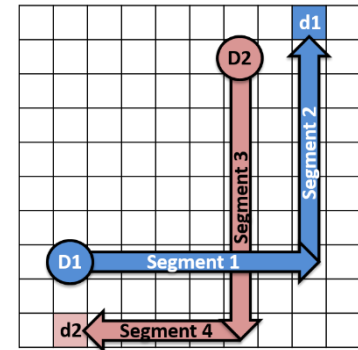
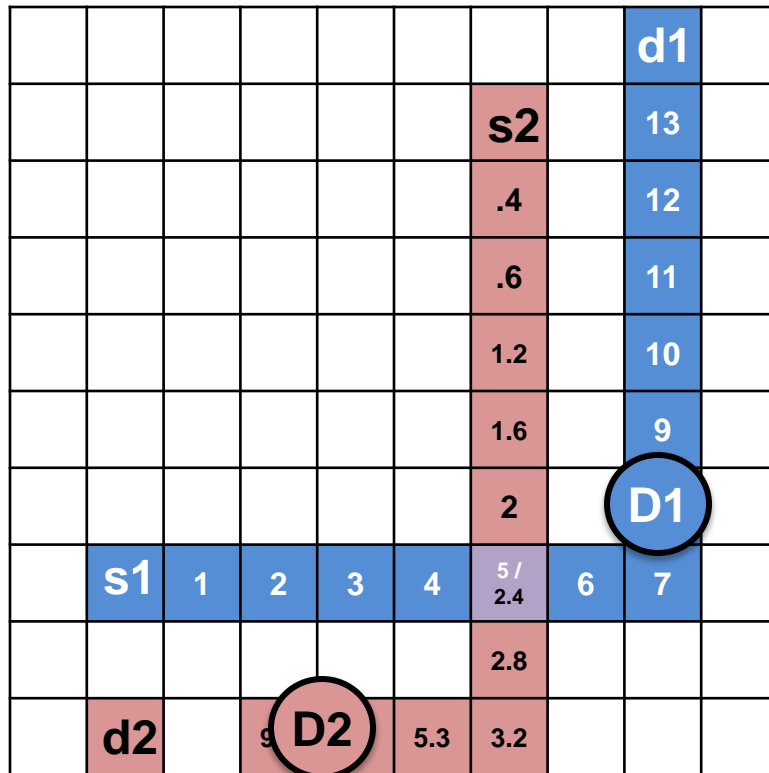


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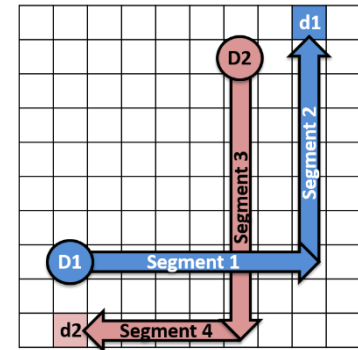
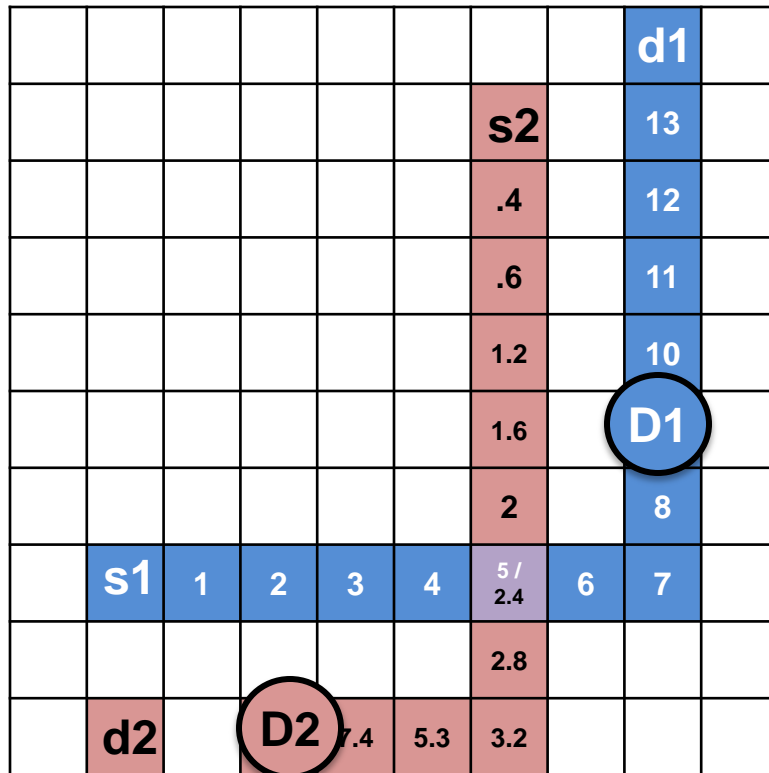


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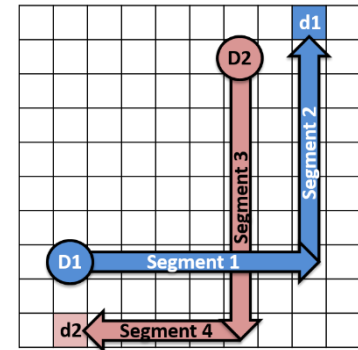
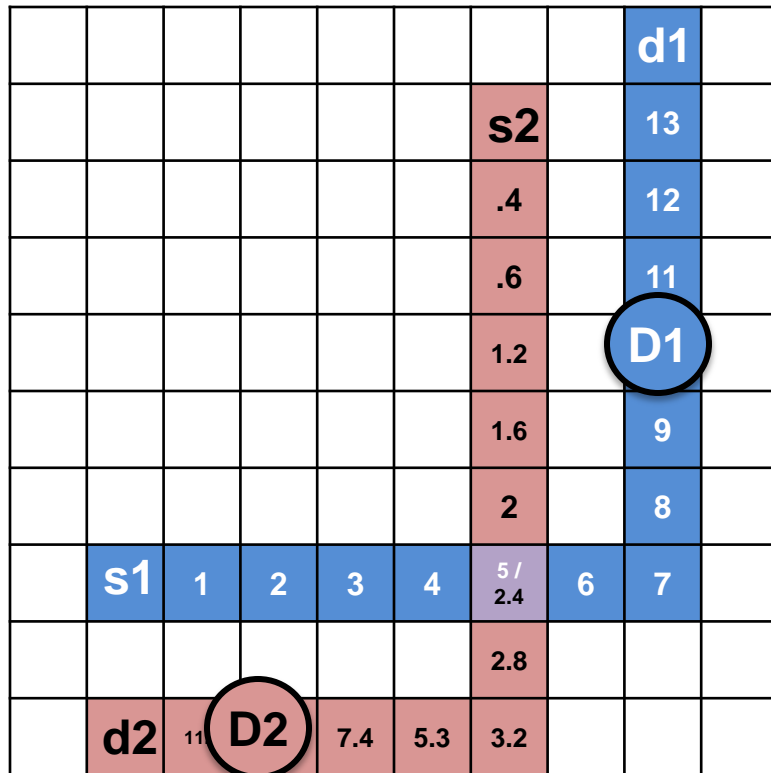


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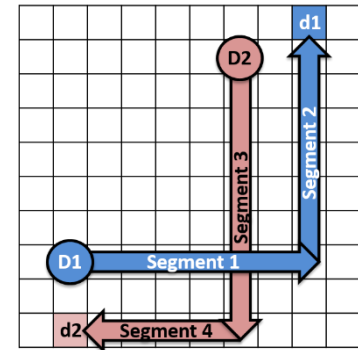
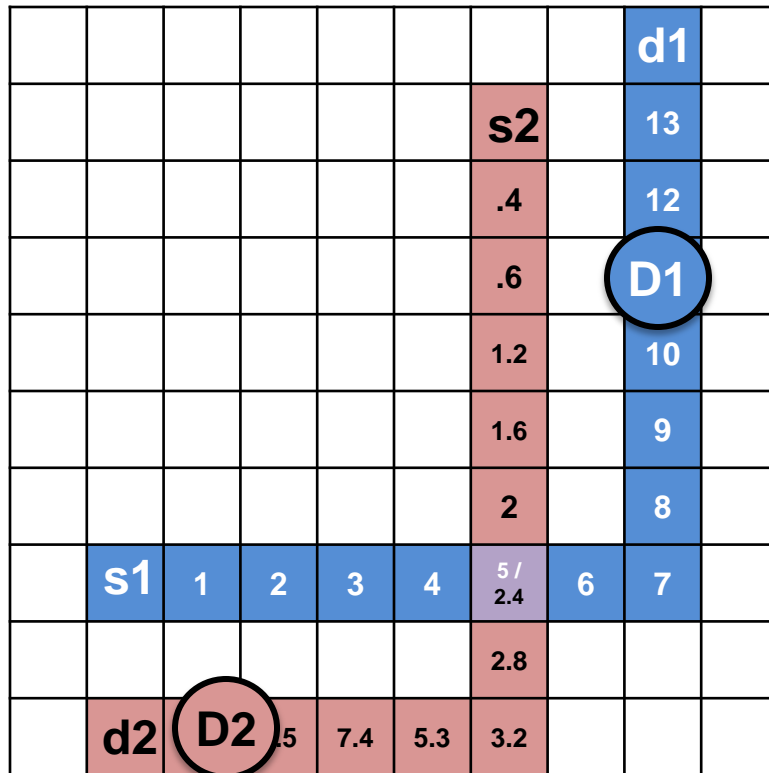


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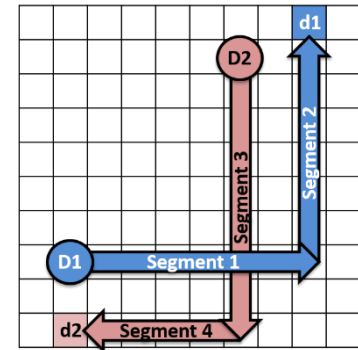
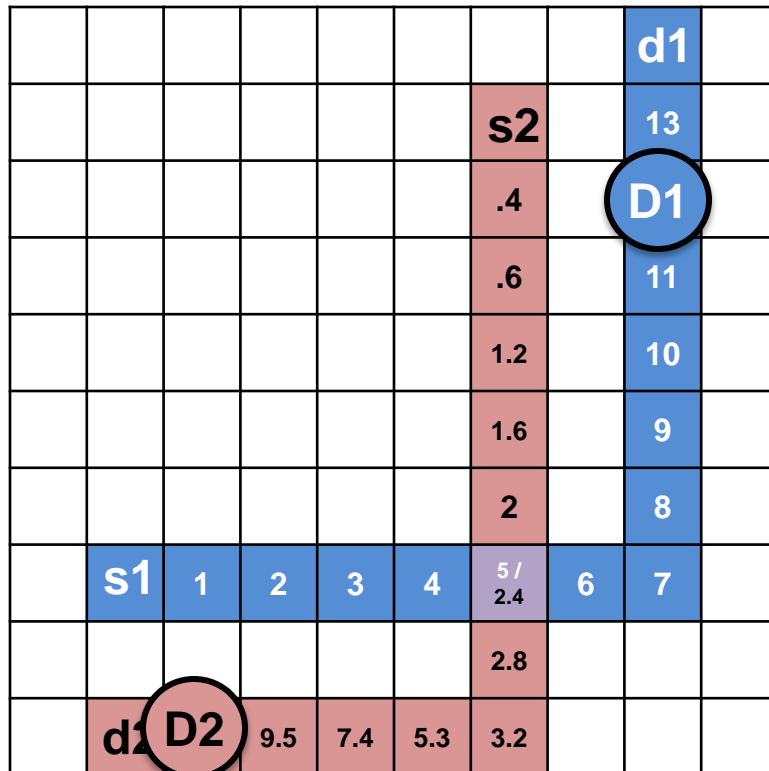


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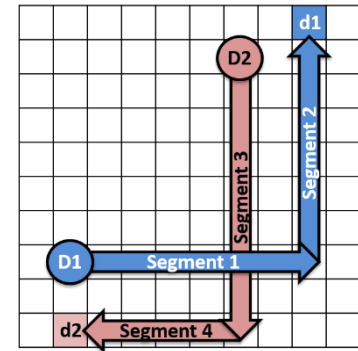
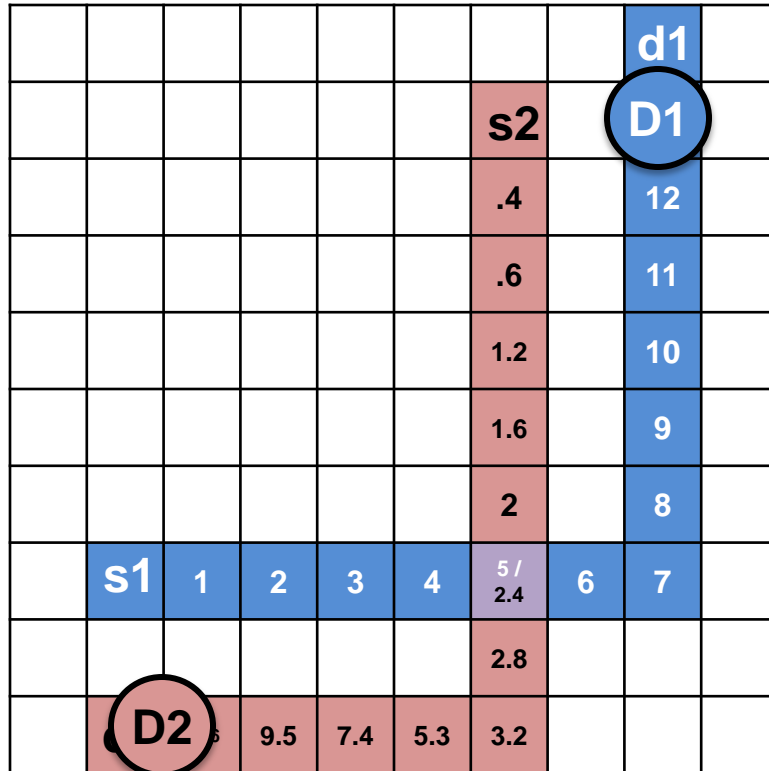


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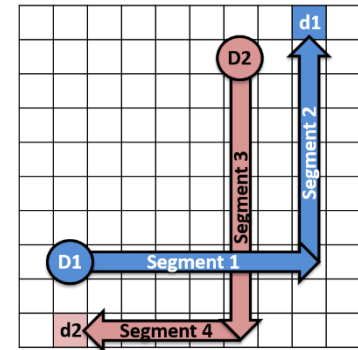
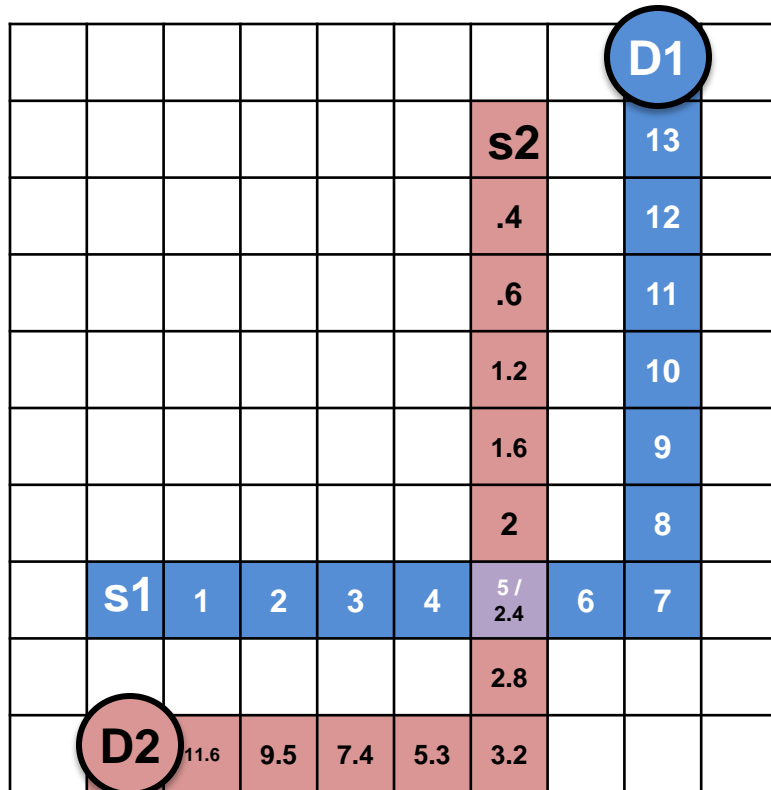


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Simple Example



Numbers on electrodes indicate the time the droplet arrives at the electrode.

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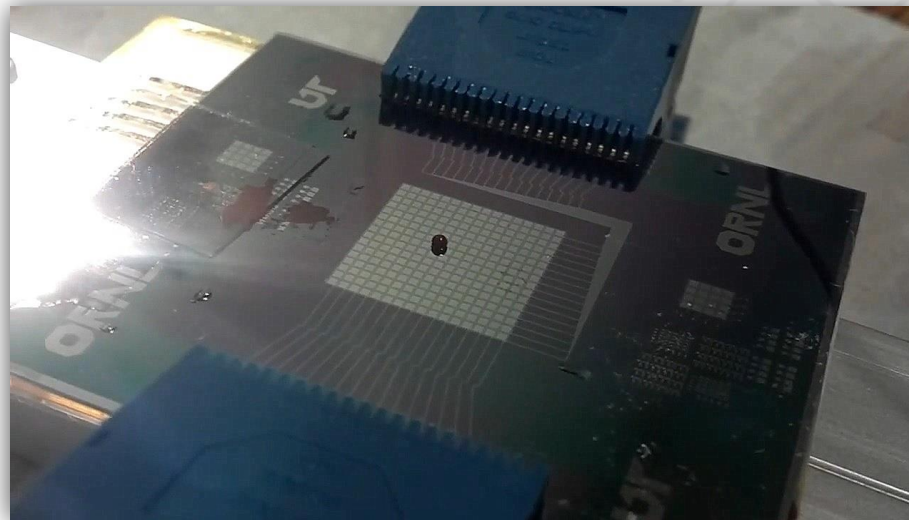
D2 compacted against D2 with no interference.

Simulation Details

- ▶ DMFB modeled after University of Tennessee's active matrix design¹

- ▶ Electrode resistance = 1GΩ
- ▶ Electrode pitch (dimension) = 2.54mm
- ▶ Voltage_{min} = 13V, Voltage_{max} = 70V

- ▶ Voltage/velocity relationship: $Velocity = 0.005 \times Voltage^2 + 0.0358 \times Voltage - 0.9103$



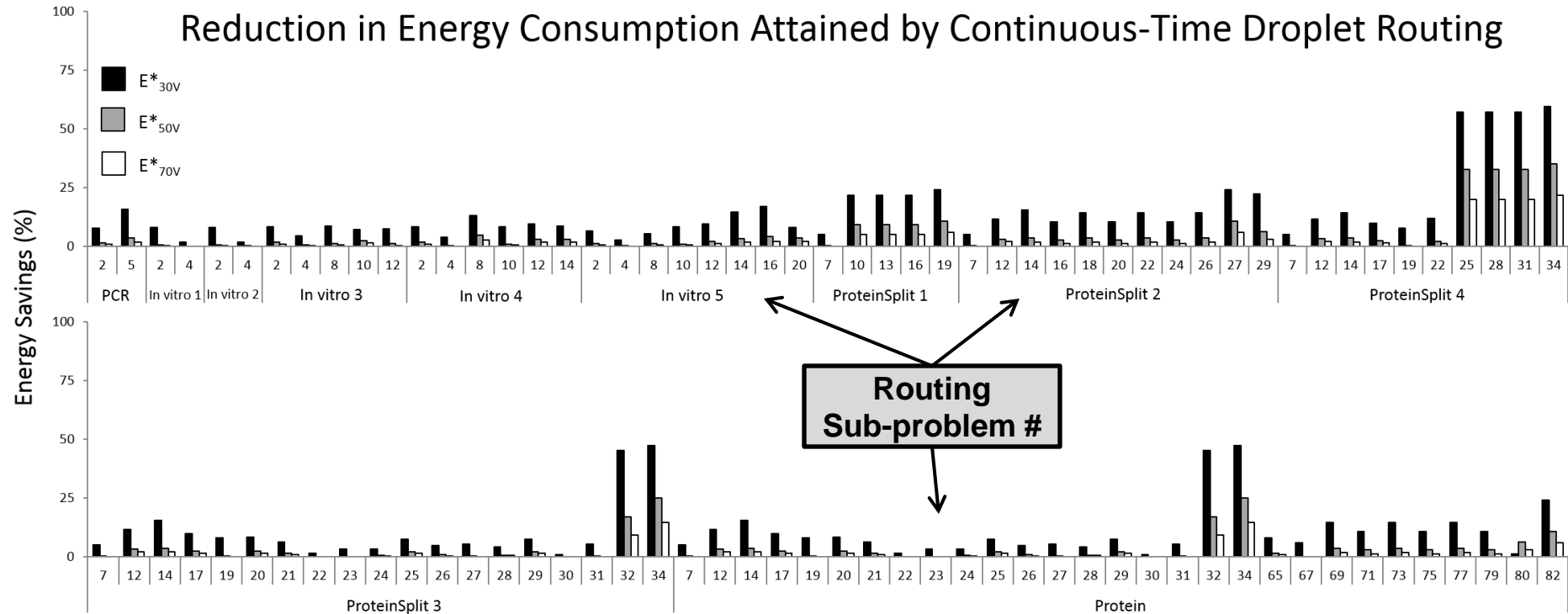
Simulation Details

- › Benchmarks
 - › PCR, In-Vitro Diagnostics, Protein, ProteinSplit assays (common benchmarks)
- › Base Routing Flow²
 - › Step 1: Roy maze router (same as proposed)
 - › Step 2: Constant voltage
 - › Add stalls at beginning of routes to avoid interference
- › Setup
 - › Schedules and placements same for both route compactors

² Grissom, D., and Brisk, P. Fast online synthesis of generally programmable digital microfluidic biochips. In *Proceedings of the ACM/IEEE International Conference on Hardware Software Codesign and System Synthesis* (Tampere, Finland, October 07 - 12, 2012). CODES-ISSS '12, 413-422.

Results: Energy Savings

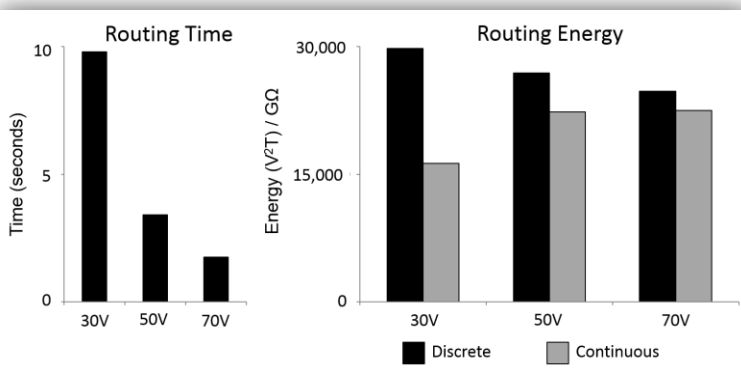
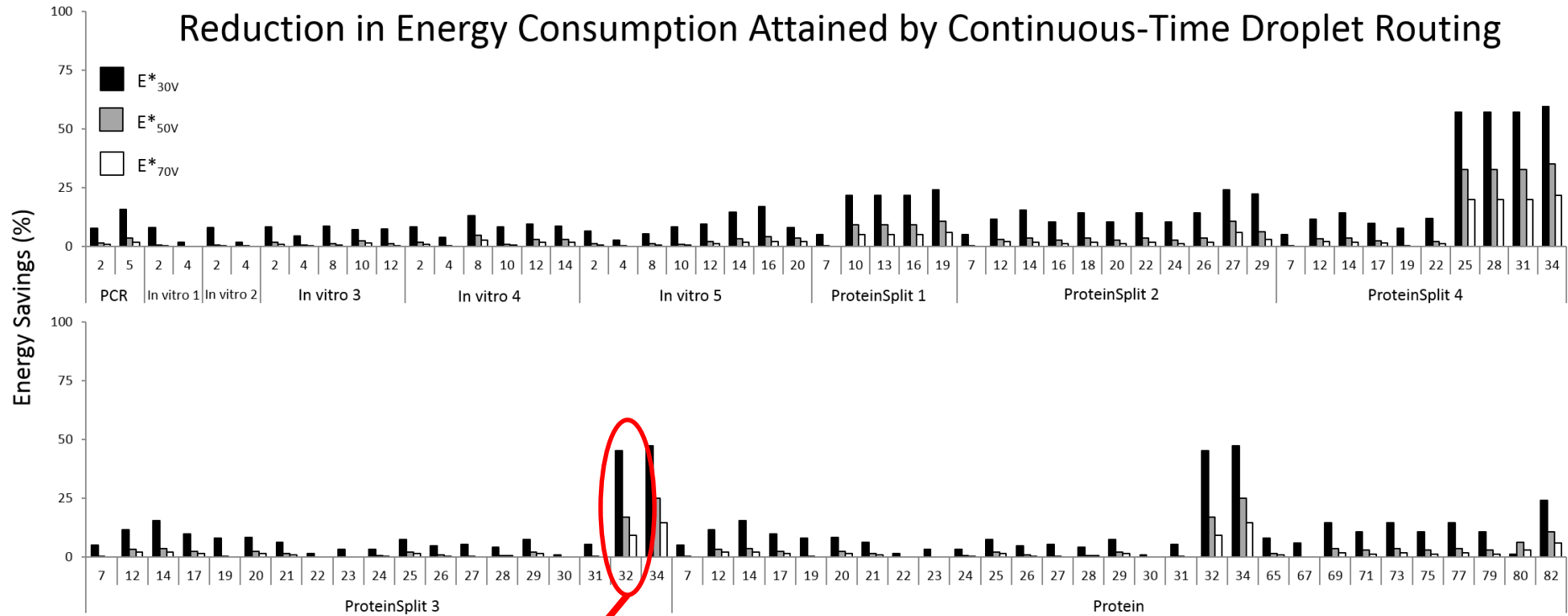
Reduction in Energy Consumption Attained by Continuous-Time Droplet Routing



- ▶ Base flow performed at 30V, 50V and 70V
 - ▶ Time constraints for continuous-time compaction derived from these runs
- ▶ Energy savings vary greatly between sub-problems
 - ▶ Due to amount and complexity of droplets being routed

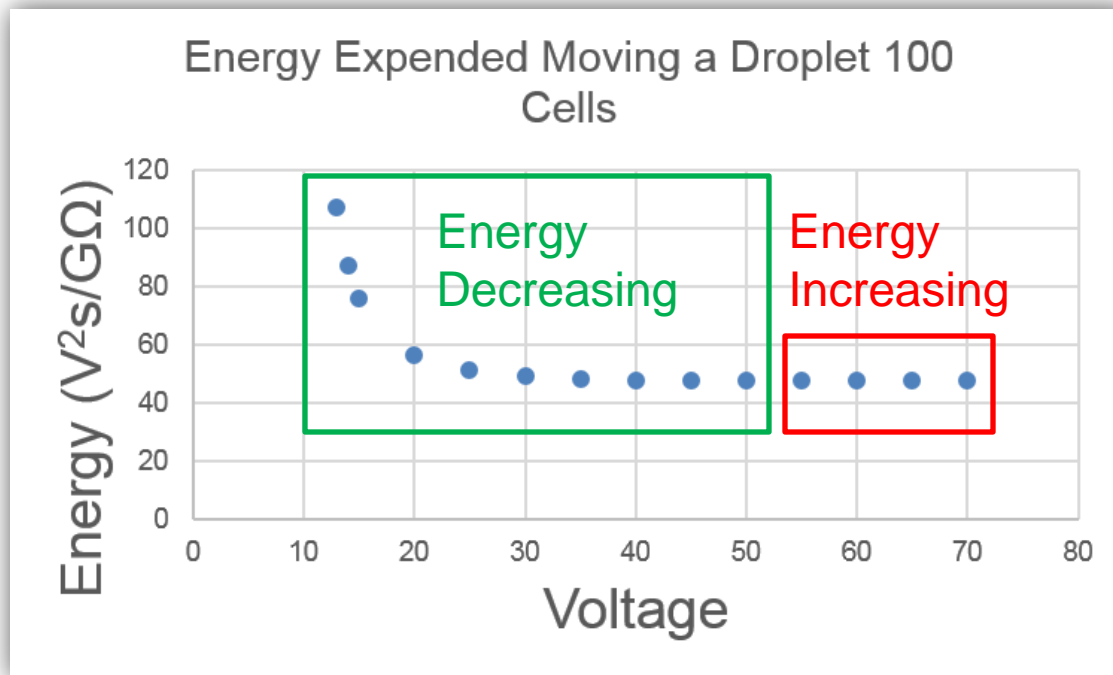
Results: Energy Savings

Reduction in Energy Consumption Attained by Continuous-Time Droplet Routing



- Higher voltages → Better energy usage across platforms
- More V for less time can lead to energy savings
- 30V sees greatest savings because slower paths provide more opportunities for route speedups

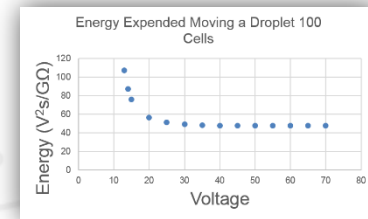
Results: Energy Savings



- ▶ Threshold exists where *Increasing Voltage* → *Decreases Energy* becomes not true
- ▶ Threshold depends on device characteristics
- ▶ Large savings can be incurred by decreasing voltage on halts
 - ▶ Wait for 0.5s:
 - ▶ @ 30V → 450 V²s/GΩ
 - ▶ @ 70V → 2450 V²s/GΩ

Conclusion

- ▶ First model for continuous-time domain droplet routing (compaction)
 - ▶ Varying voltage → varying velocity
- ▶ Multiple speeds allow for energy savings
 - ▶ Higher voltages can have better energy usage
 - ▶ Continuous-time domain droplet compaction can achieve energy savings across range of voltage
- ▶ Tradeoffs may vary based on characteristics of DMFB



Thank You

<http://microfluidics.cs.ucr.edu/>

