

A Two-Stage ILP-Based Droplet Routing Algorithm For Pin-Constrained Digital Microfluidic Biochips

**2010 ACM International Symposium on Physical Design
(ISPD'10)**

Tsung-Wei Huang and Tsung-Yi Ho

<http://eda.csie.ncku.edu.tw>

Department of Computer Science and Information Engineering
National Cheng Kung University
Tainan, Taiwan



Outline

- . Introduction
- . Problem formulation
- . Our contribution
- . Basic ILP formulation
- . Deterministic ILP formulation
- . Experimental results
- . Conclusion



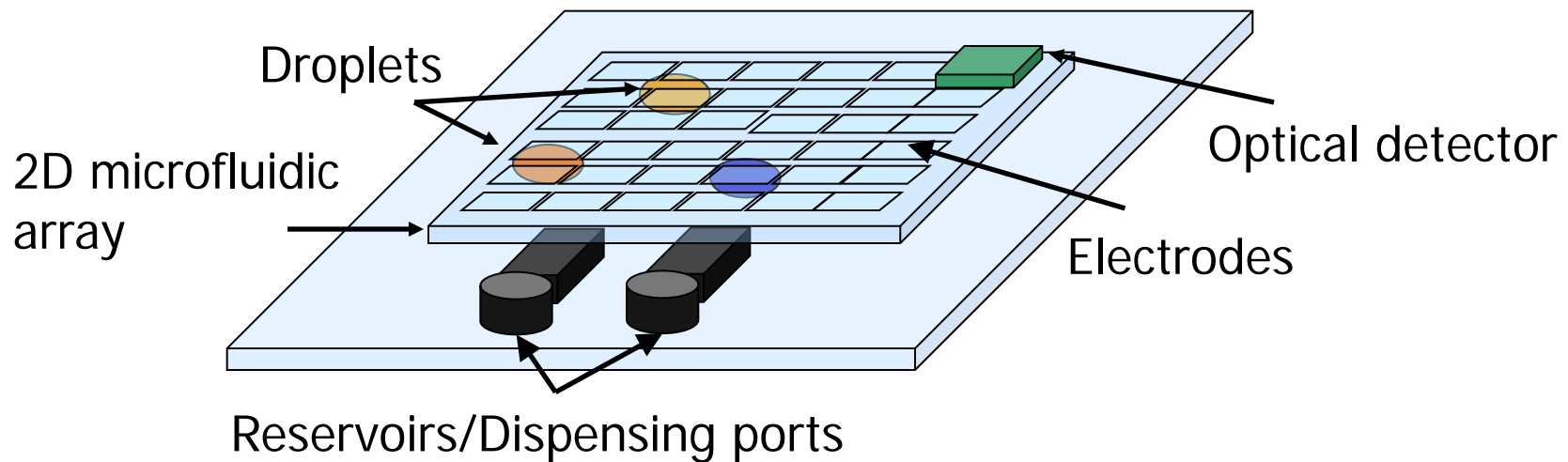
Outline

- . Introduction
 - Digital microfluidic biochips
 - Pin-constrained digital microfluidic biochips
 - Previous work and limitations
- . Our contribution
- . Problem formulation
- . Basic ILP formulation
- . Deterministic ILP formulation
- . Experimental results
- . Conclusion



Digital Microfluidic Biochips (DMFBs) (1/2)

- Three main components:
 - **2D microfluidic array:** set of basic cells for biological reactions
 - **Reservoirs/dispensing ports:** for droplet generation
 - **Optical detectors:** detection of reaction result
- Perform laboratory procedures based on *droplets*
 - **Droplet:** biological sample carrier

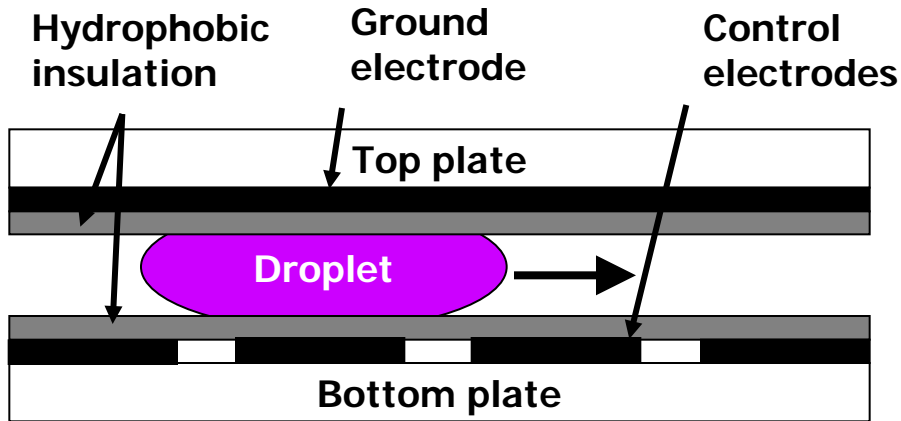


The schematic view of a biochip (Duke Univ.)

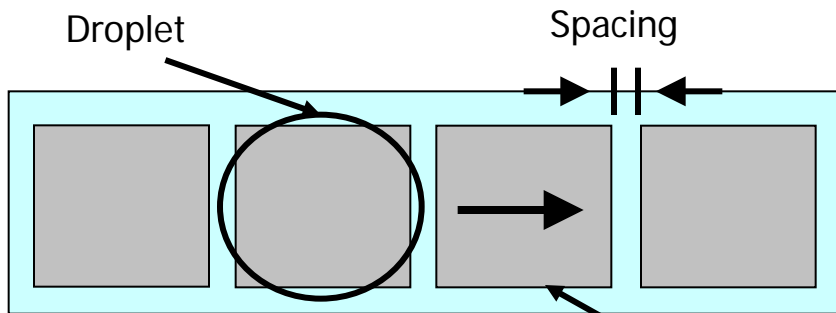


Digital Microfluidic Biochips (DMFBs) (2/2)

- Movement control of a droplet

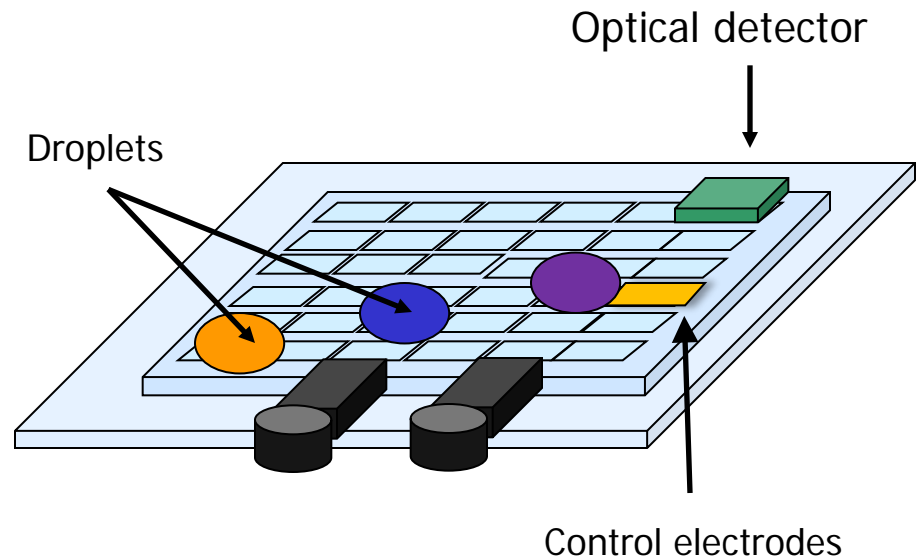


Side view



Top view

Generated electrical force

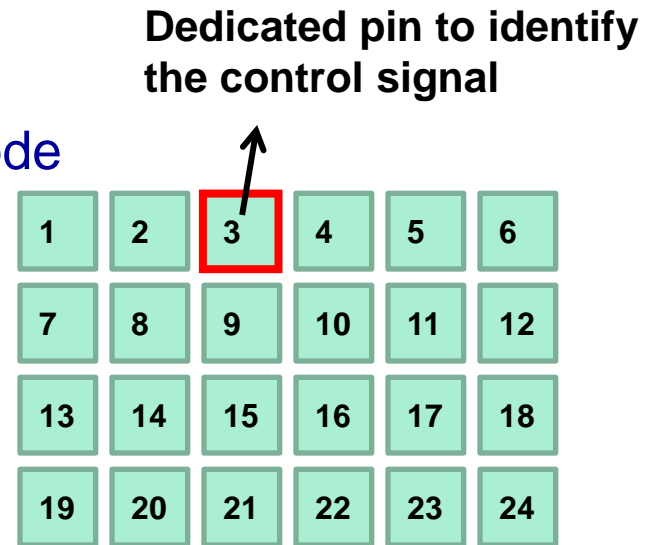


Pin-Constrained Digital Microfluidic Biochips

Direct-addressing biochips

- Dedicated control pin for each electrode
- Maximum freedom of droplets
- High demanded control pins

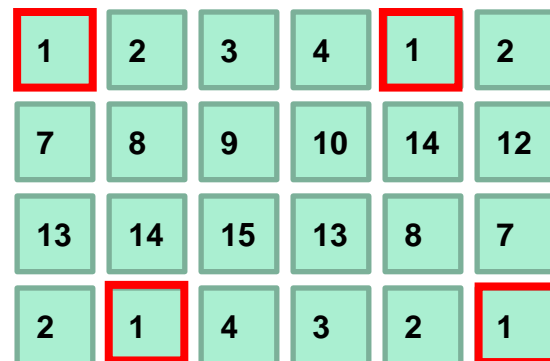
Control pins: 24



Broadcast-addressing biochips *

- A control pin can be shared by multiple electrodes
- Flexible for pin-constrained DMFBs
- Control pin sharing

Control pins: 15



* [T. Xu and K. Chakrabarty, DAC'08]



Previous Work and Limitation (1/2)

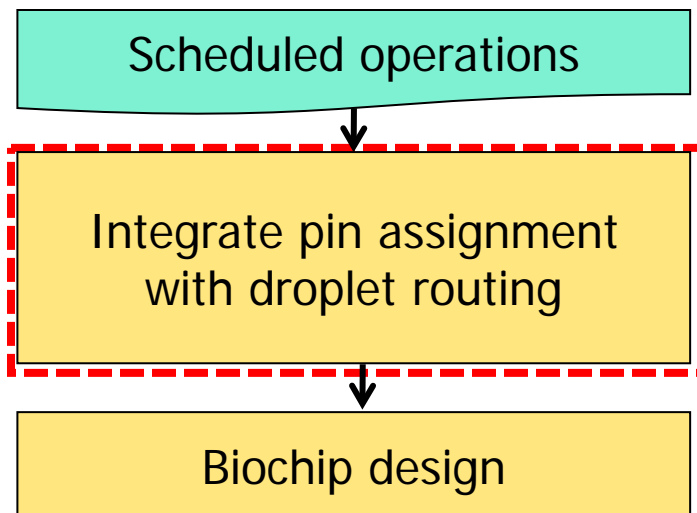
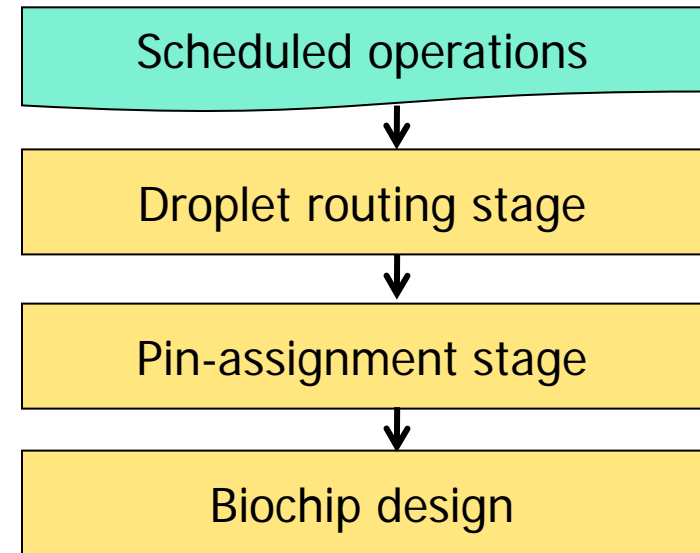
- Droplet routing algorithms
 - Droplet routing in the synthesis of digital microfluidic biochips
[Su et al, DATE'06]
 - Modeling and controlling parallel tasks in droplet based microfluidic systems
[K. F. Böhringer, TCAD'06]
 - A network-flow based routing algorithm for digital microfluidic biochips
[Yuh et al, ICCAD'07]
 - Integrated droplet routing in the synthesis of microfluidic biochips
[T. Xu and K. Chakrabarty, DAC'07]
 - A high-performance droplet routing algorithm for digital microfluidic biochips
[Cho and Pan, ISPD'08]
- Pin-constrained digital microfluidic biochips
 - Droplet-trace-based array partition and a pin assignment algorithm for the automated design of digital microfluidic biochips
[T. Xu and K. Chakrabarty, CODES+ISSS'06]
 - Broadcast electrode-addressing for pin-constrained multi-functional digital microfluidic biochips
[T. Xu and K. Chakrabarty, DAC'08]



Previous Work and Limitation (2/2)

. Limitations

- Separately consider the routing stage and the pin-assignment stage
- The solution quality is limited
 - # of Control pins
 - # of Used cells
 - Execution time



Ours integrated method **simultaneously** minimizes the # of control pins, # of used cells, and execution time for pin-constrained DMFBs.



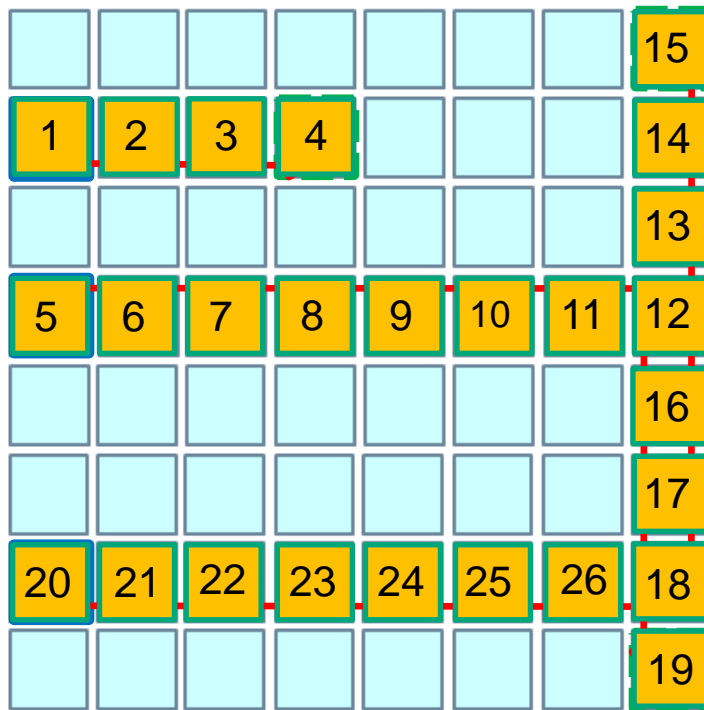
Outline

- . Introduction
- . **Our contribution**
- . Problem formulation
- . Basic ILP formulation
- . Deterministic ILP formulation
- . Experimental results
- . Conclusion



Previous Method – Direct Addressing

- Apply the direct addressing to a routing result
 - Separate pin assignment stage and routing stage



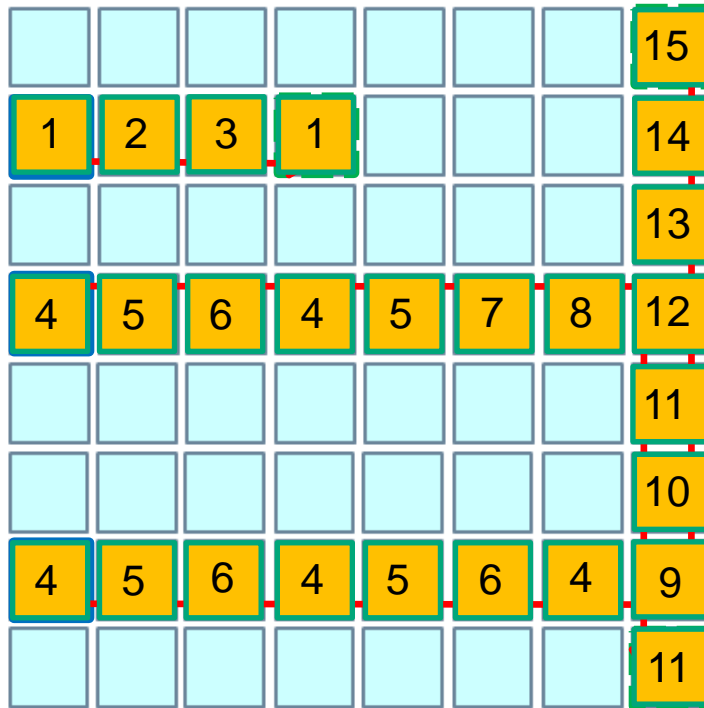
Control Pins: 26
Used Cell: 26
execution time: 18

of control pins = # of used cells



Previous Method (1/2) – Broadcast Addressing

- Apply the broadcast addressing to a routing result
 - Separate pin assignment stage and routing stage

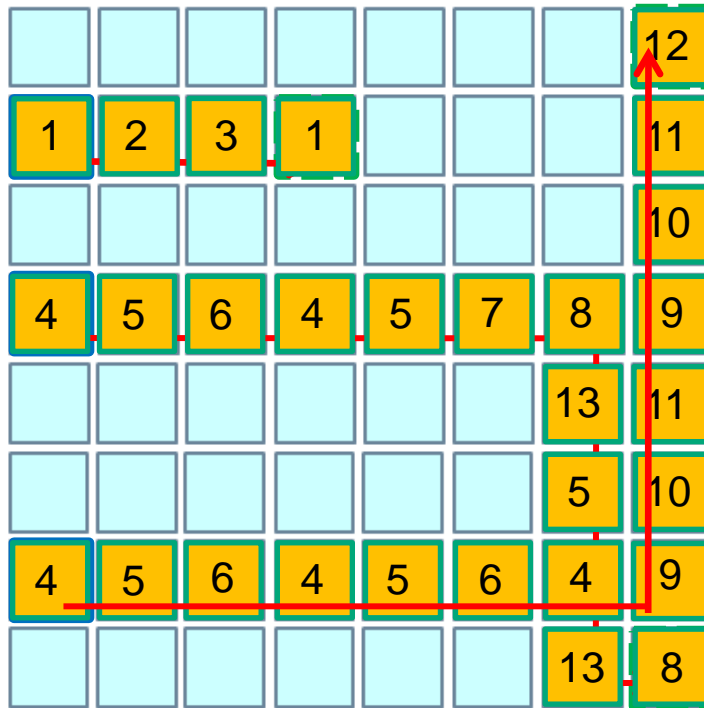


Control Pins: 15
Used Cell: 26
execution time: 18



Previous Method (2/2) – Broadcast Addressing

- Simply **integrate** the broadcast addressing with droplet routing



Control Pins: 15
Used Cell: 26
execution time: 18

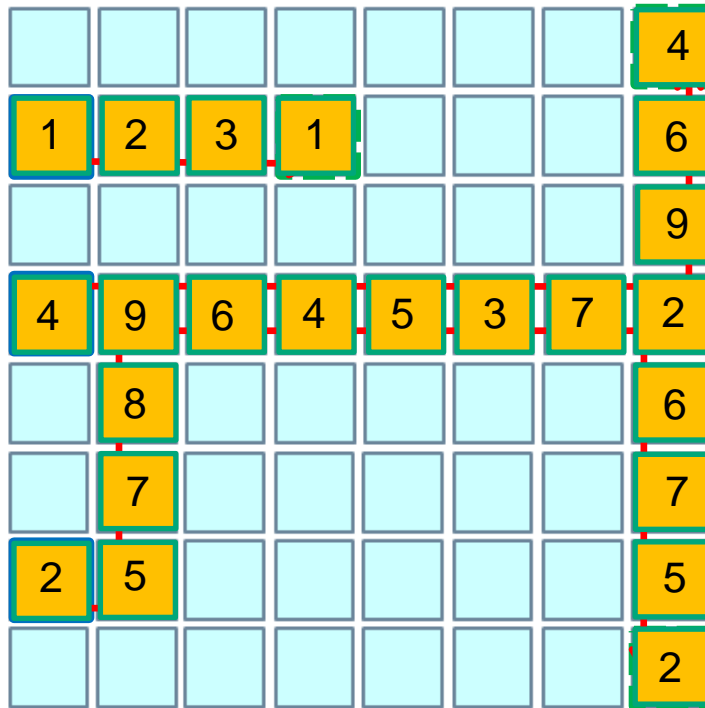
Control Pins: 13
Used Cell: 29
execution time: 20

May increase the # of used cells
and execution time



Ours (1/2)

- Integrate broadcast addressing with droplet routing while **simultaneously** minimizing the # of control pins, # of used cells, and execution time



Control Pins: 15
Used Cell: 26
execution time: 18

Control Pins: 13
Used Cell: 29
execution time: 20

Control Pins: 9
Used Cell: 23
execution time: 15

Minimized # of control pins
Minimized # of used cells
Minimized execution time



Ours (2/2)

. Contributions:

- We propose the first algorithm that integrates the broadcast-addressing with droplet routing problem, while **simultaneously** minimizing the # of control pins, # of used cells, and execution time
- A basic ILP formulation is introduced to obtain an **optimal solution**
- A two-stage ILP-based algorithm is presented to tackle the complexity of the basic ILP formulation



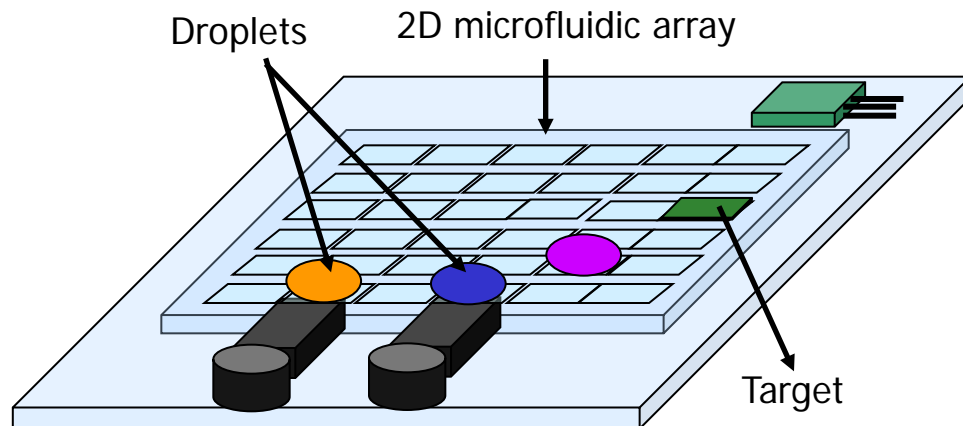
Outline

- . Introduction
- . Our contribution
- . **Problem formulation**
- . Basic ILP formulation
- . Deterministic ILP formulation
- . Experimental results
- . Conclusion

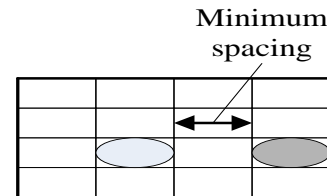


Problem Formulation

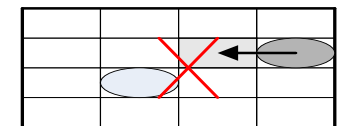
- **Input:** A netlist of n droplets $D = \{d_1, d_2, \dots, d_n\}$, the locations of modules
- **Objective:** Route all droplets from their source cells to their target cells while *minimizing the # of control pins, # of used cells, and execution time* for high throughput designs
- **Constraint:** Fluidic and timing constraints should be satisfied.



- **Fluidic constraint**



Static fluidic constraint



Dynamic fluidic constraint

- **Timing constraint**

- **Maximum available executed time**



Outline

- . Introduction
- . Problem formulation
- . Our contribution
- . Basic ILP formulation
 - Objective function
 - Basic constraints
 - Electrode constraints
 - Broadcast-addressing constraints
 - Limitations
- . Deterministic ILP formulation
- . Experimental results
- . Conclusion



Objective Function

- Objective function
 - Minimize the # of control pins (product cost)
 - Minimize the # of used cells (fault-tolerance)
 - Minimize the execution time (reliability)

$$\text{Minimize : } \alpha \sum up(p) + \beta \sum uc(x, y) + \gamma T_l$$

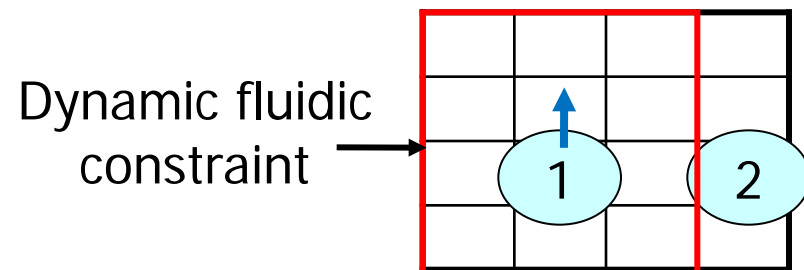
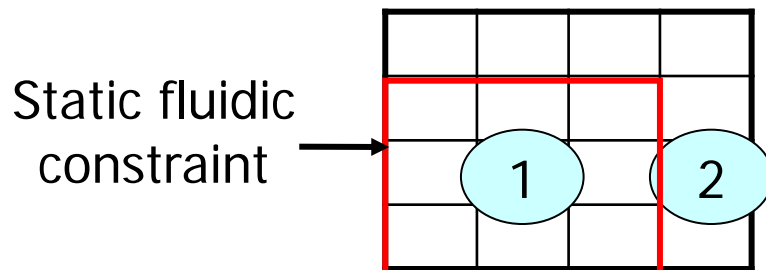
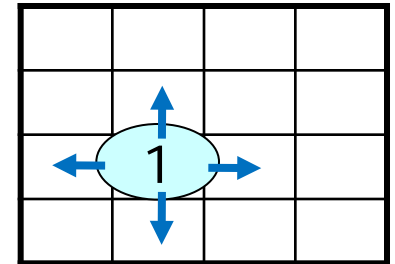
α # of control pins β # of used cells γ execution time

where α , β , and γ are user-defined parameters



Basic Constraints

- Source/target requirement
 - All droplets locate at their sources at time zero
 - A droplet stays at its target once reaching it
- Exclusive constraint
 - Each droplet has only one location at a time step
- Droplet movement constraint
 - A droplet can move to four adjacent cells or stall
- Static/dynamic fluidic constraint
 - No other droplets are in the 3x3 region centered by a droplet **at time t / within $t \sim t+1$**



Electrode Constraints (1/2)

- . Electrode constraints
 - To model the control of droplets by turning on/off the actuation voltage of electrodes
- . Activation type
 - “1” represents the activated electrode (turn on)
 - “0” represents the deactivated electrode (turn off)
 - “X” represents the don’t care (both “1” and “0” are legal)
- . Formulation technique
 - Extract the cells that “**must-be-activated**”
 - Extract the cells that “**must-be-deactivated**”



Electrode Constraints (2/2)

Illustration

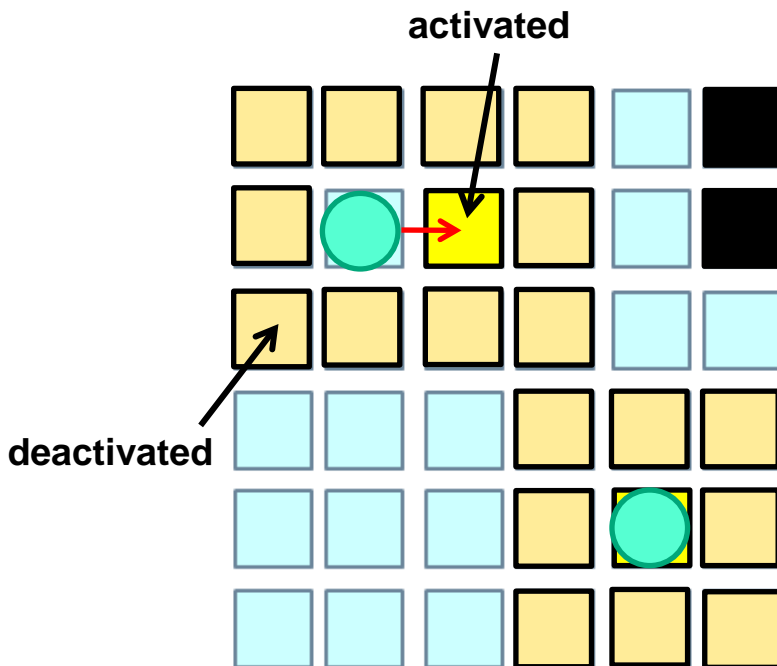
 Must be deactivated(0)

 Must be activated (1)

 Blockage

 Droplet

 Don't care (X)



of activated cells: 1

of deactivated cells: 11

↓

| | | | | | |
|---|---|---|---|---|---|
| 0 | 0 | 0 | 0 | X | |
| 0 | 0 | 1 | 0 | X | |
| 0 | 0 | 0 | 0 | X | X |
| X | X | X | 0 | 0 | 0 |
| X | X | X | 0 | 1 | 0 |
| X | X | X | 0 | 0 | 0 |

of activated cells: 1

of deactivated cells: 8



Broadcast-Addressing Constraints

- Broadcast-addressing constraints
 - Model the pin assignment by “compatible” activation sequences

| Electrode | E_1 | E_2 | E_3 | E_4 | E_5 | E_6 | E_7 | E_8 | E_9 | E_{10} | E_{11} | E_{12} |
|---------------------|-------|-------|-------|-------|-------|-------|-------|-------|-------|----------|----------|----------|
| Activation sequence | 1 | 1 | 0 | 0 | 0 | X | X | 0 | X | X | X | X |
| | 0 | 0 | 1 | 1 | 1 | 0 | 0 | 1 | X | X | X | X |
| | 0 | 0 | 0 | 0 | 0 | 1 | 1 | 0 | 0 | 0 | X | X |
| | X | X | 0 | 0 | 0 | 0 | 0 | 0 | 1 | 1 | 0 | 0 |
| | X | X | X | X | 1 | 0 | 0 | 1 | X | X | 1 | 1 |

Merge: E_4 and E_5 $0100X+01001 \rightarrow 01001$

| Pin-assignment result | | |
|-----------------------|------------|----------------------------|
| Pin | Electrodes | Merged activation sequence |
| 1 | E_4, E_5 | 0 1 0 0 1 |

or

| Pin-assignment result | | |
|-----------------------|------------|----------------------------|
| Pin | Electrodes | Merged activation sequence |
| 1 | E_4 | 0 1 0 0 X |
| 2 | E_5 | 0 1 0 0 1 |

Merge: E_5 and E_6 $01001+X0100 \rightarrow$ **Invalid**



Limitations

- . Pros and cons
 - Advantage: ***an optimal solution***
 - Drawback: ***only feasible to small applications***
- . Multi-objectives optimization
 - Simultaneously consider the optimization of the #of control pins, # of used cells, and execution time
 - Introduce a high solution space
- . Many formulation constraints
 - High # of variables
 - High # of constraints



Outline

- . Introduction
- . Problem formulation
- . Our contribution
- . Basic ILP formulation
- . **Deterministic ILP formulation**
 - Two-stage ILP-based routing algorithm
- . Experimental results
- . Conclusion



Two-Stage ILP-Based Routing Algorithm

- . First stage
 - Major goal: **reduce the solution space**
 - Global routing
 - Obtain an initial routing paths

- . Second stage
 - Major goal: **accelerate the searching time**
 - Incremental ILP-based routing method
 - Iteratively select an un-routed droplet
 - Route this droplet with previous routing solutions



Global Routing

Global routing

- Preferred routing tracks construction
 - Reduce the design complexity
- A* maze search for min-cost routing path
 - Orderly routing along these tracks
 - Minimized used cells



Source location



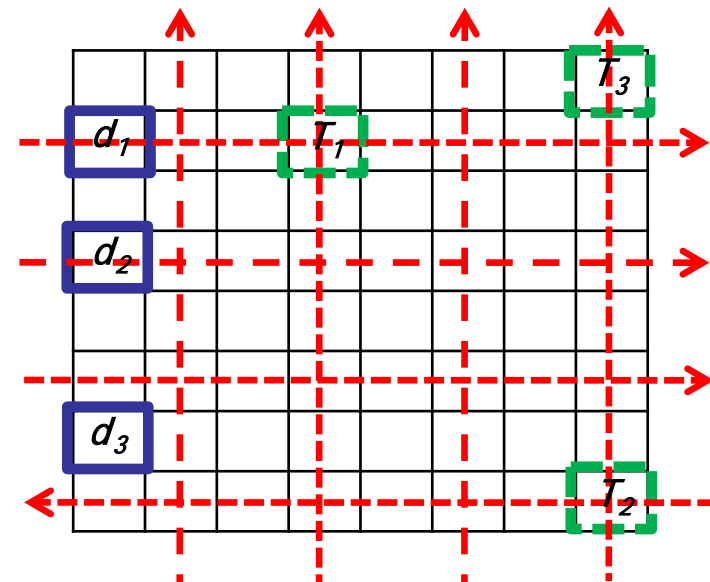
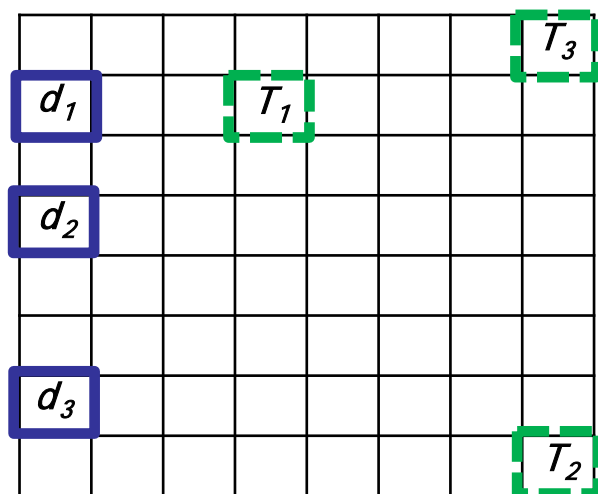
Global routing track



Sink location



Updated global routing track



T.-W. Huang, C.-H. Lin and T.-Y. Ho, " A Contamination Aware Droplet Routing Algorithm for Digital Microfluidic Biochips," *Proceedings of ACM/IEEE ICCAD 2009*



Incremental ILP-Based Routing (1/3)

- Net criticality calculation
 - Determine the routing order globally
 - Consider the interferences and congestion issue between droplets
 - A droplet d_i is said to be critical if d_i has fewer possible routing solutions

$$\text{crit}(d_i) = \frac{(|E_b^i| + |E_s^i|) - |E_t^i|}{|BB_i|}$$

$$E_b^i = \{c \mid c \in E_b \cap BB_i\}$$

$$E_s^i = \{c \mid c \in E_{s_j} \cap BB_i, \forall d_j \in D / d_i\}$$

$$E_t^i = \{c \mid c \in E_{t_j} \cap BB_i, \forall d_j \in D / d_i\}$$



Incremental ILP-Based Routing (2/3)

. Deterministic ILP

- Select an un-routed droplet
- Routing resources: M_i
 - Maximum available routing time T_l^i
 - Maximum available control pins P_l^i
- Increasing scalar: IS
 - Growth rate of routing resources

$$M_i = (T_l^i + \sigma_1 IS) + (P_l^i + \sigma_2 IS)$$

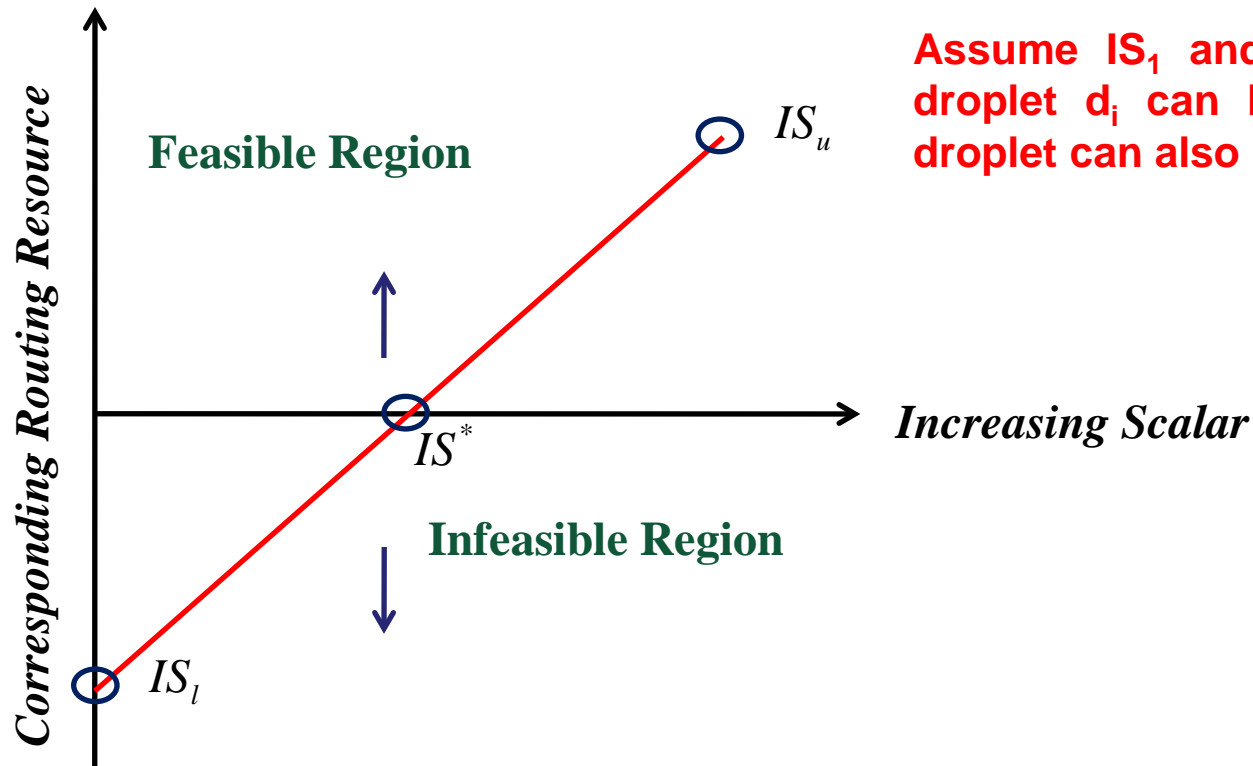
- Major goal:
 - Determine the feasibility with the given routing resources
 - Objective function:

Minimize : c



Incremental ILP-Based Routing (3/3)

- Monotonic property
 - Binary solution search method
 - Logarithmic number of searching iterations



Outline

- . Introduction
- . Problem formulation
- . Our contribution
- . Basic ILP formulation
- . Deterministic ILP formulation
- . [Experimental results](#)
- . Conclusion



Experimental Results (1/5)

- Implement our algorithm in C++ language on a 2 GHz 64-bit Linux machine with 16GB memory
- Compare with
 - Network flow algorithm [P.-H Yuh et al, ICCAD'07]
 - High performance [M. Cho and D. Z. Pan, TCAD'08]
- Statistic of benchmarks

| Benchmark | Size | #Sub | T_{\max} | #Nets | #D _{max} |
|-----------|---------|------|------------|-------|-------------------|
| vitro_1 | 16 X 16 | 11 | 20 | 28 | 5 |
| vitro_2 | 14 X 14 | 15 | 20 | 35 | 6 |
| protein_1 | 21 X 21 | 64 | 20 | 181 | 6 |
| protein_2 | 13 X 13 | 78 | 20 | 178 | 6 |

■ Size: size of microfluidic array. ■ #Sub: # of subproblems. ■ T_{\max} : timing constraint.
■ #Nets: total # of nets. ■ #D_{max}: maximum # of droplets among subproblems.



Experimental Results (2/5)

Comparison of the # of control pins

| Benchmark | Direct Addressing | | Broadcast Addressing | | Two-Stage ILP | | |
|-----------|-------------------|-------------|----------------------|-------------|---------------|-------------|-----------|
| | [11] | [4] | [11]+[10] | [4]+[10] | [11]+IILP | [4]+IILP | Ours |
| | P_{avg} | P_{avg} | P_{avg} | P_{avg} | P_{avg} | P_{avg} | P_{avg} |
| vitro_1 | 21.55 | 23.45 | 9.48 | 10.11 | 9.11 | 9.49 | 4.51 |
| vitro_2 | 15.73 | 16.40 | 8.95 | 10.64 | 8.03 | 9.21 | 5.01 |
| protein_1 | 25.28 | 26.38 | 9.52 | 10.55 | 8.54 | 9.25 | 5.43 |
| protein_2 | 12.03 | 12.35 | 8.73 | 8.55 | 7.72 | 7.38 | 4.43 |
| | 3.82 | 4.03 | 1.90 | 2.06 | 1.73 | 1.83 | 1 |

[4] M. Cho and D. Z. Pan, "A high-performance droplet routing algorithm for digital microfluidic biochips," IEEE Trans. on CAD, vol. 27, no. 10, pp. 1714-1724, Oct. 2008.

[10] T. Xu and K. Chakrabarty, "Broadcast electrode-addressing for pin-constrained multi-functional digital microfluidic biochips," Proc. IEEE/ACM DAC, pp. 173-178, Jun. 2008.

[11] P.-H. Yuh, C.-L. Yang, and Y.-W. Chang, "BioRoute: A network-flow based routing algorithm for digital microfluidic biochips," Proc. IEEE/ACM ICCAD, pp. 752-757, Nov. 2007.



Experimental Results (3/5)

. Comparison of the # of used cells

| Benchmark | Direct Addressing | | Broadcast Addressing | | Two-Stage ILP | | |
|-----------|-------------------|------|----------------------|----------|---------------|----------|------|
| | [11] | [4] | [11]+[10] | [4]+[10] | [11]+IILP | [4]+IILP | Ours |
| | U.C. | U.C. | U.C. | U.C. | U.C. | U.C. | U.C. |
| vitro_1 | 237 | 258 | 237 | 258 | 231 | 243 | 231 |
| vitro_2 | 236 | 246 | 236 | 246 | 231 | 229 | 229 |
| protein_1 | 1618 | 1688 | 1618 | 1688 | 1597 | 1627 | 1582 |
| protein_2 | 939 | 963 | 939 | 963 | 927 | 943 | 930 |
| | 1.02 | 1.07 | 1.02 | 1.07 | 1.00 | 1.02 | 1 |

[4] M. Cho and D. Z. Pan, "A high-performance droplet routing algorithm for digital microfluidic biochips," IEEE Trans. on CAD, vol. 27, no. 10, pp. 1714-1724, Oct. 2008.

[10] T. Xu and K. Chakrabarty, "Broadcast electrode-addressing for pin-constrained multi-functional digital microfluidic biochips," Proc. IEEE/ACM DAC, pp. 173-178, Jun. 2008.

[11] P.-H. Yuh, C.-L. Yang, and Y.-W. Chang, "BioRoute: A network-flow based routing algorithm for digital microfluidic biochips," Proc. IEEE/ACM ICCAD, pp. 752-757, Nov. 2007.



Experimental Results (4/5)

Comparison of the execution time

| Benchmark | Direct Addressing | | Broadcast Addressing | | Two-Stage ILP | | |
|-----------|-------------------|-------------|----------------------|-------------|---------------|-------------|------------|
| | [11] | [4] | [11]+[10] | [4]+[10] | [11]+ILP | [4]+ILP | Ours |
| | Avg. T_i | Avg. T_i | Avg. T_i | Avg. T_i | Avg. T_i | Avg. T_i | Avg. T_i |
| vitro_1 | 13.00 | 14.30 | 13.00 | 14.30 | 12.47 | 13.55 | 12.41 |
| vitro_2 | 11.33 | 12.00 | 11.33 | 12.00 | 11.01 | 11.48 | 10.46 |
| protein_1 | 16.31 | 16.55 | 16.31 | 16.55 | 16.08 | 15.44 | 15.42 |
| protein_2 | 10.51 | 12.19 | 10.51 | 12.19 | 10.33 | 11.52 | 10.22 |
| | 1.05 | 1.14 | 1.05 | 1.14 | 1.03 | 1.08 | 1 |

[4] M. Cho and D. Z. Pan, "A high-performance droplet routing algorithm for digital microfluidic biochips," IEEE Trans. on CAD, vol. 27, no. 10, pp. 1714-1724, Oct. 2008.

[10] T. Xu and K. Chakrabarty, "Broadcast electrode-addressing for pin-constrained multi-functional digital microfluidic biochips," Proc. IEEE/ACM DAC, pp. 173-178, Jun. 2008.

[11] P.-H. Yuh, C.-L. Yang, and Y.-W. Chang, "BioRoute: A network-flow based routing algorithm for digital microfluidic biochips," Proc. IEEE/ACM ICCAD, pp. 752-757, Nov. 2007.



Experimental Results (5/5)

. Comparison of the runtime

| Benchmark | Basic ILP | [11]+IILP | [4]+IILP | Ours |
|-----------|-------------|-------------|-------------|-----------|
| | CPU (min) | CPU (sec) | CPU (sec) | CPU (sec) |
| vitro_1 | > 7200 | 14.33 | 15.31 | 10.11 |
| vitro_2 | > 7200 | 16.49 | 18.38 | 8.32 |
| protein_1 | > 7200 | 28.43 | 34.51 | 30.13 |
| protein_2 | > 7200 | 22.16 | 28.33 | 21.38 |
| | N.C. | 1.34 | 1.55 | 1 |

[4] M. Cho and D. Z. Pan, "A high-performance droplet routing algorithm for digital microfluidic biochips," IEEE Trans. on CAD, vol. 27, no. 10, pp. 1714-1724, Oct. 2008.

[10] T. Xu and K. Chakrabarty, "Broadcast electrode-addressing for pin-constrained multi-functional digital microfluidic biochips," Proc. IEEE/ACM DAC, pp. 173-178, Jun. 2008.

[11] P.-H. Yuh, C.-L. Yang, and Y.-W. Chang, "BioRoute: A network-flow based routing algorithm for digital microfluidic biochips," Proc. IEEE/ACM ICCAD, pp. 752-757, Nov. 2007.



Outline

- . Introduction
- . Problem formulation
- . Our contribution
- . Basic ILP formulation
- . Deterministic ILP formulation
- . Experimental results
- . Conclusion



Conclusion

- . We proposed the first algorithm that integrates the broadcast-addressing with the droplet routing problem while simultaneously minimizing the # of control pins, # of used cells, and execution time
- . A basic ILP formulation is introduced to optimally solve this problem
- . A two-stage ILP-based routing algorithm is also presented to tackle the complexity of the basic ILP formulation
- . Experimental results demonstrate that our algorithm achieves the best results in terms of the # of control pins, # of used cells, and execution time.



**Thank You for
Your Attention!**

