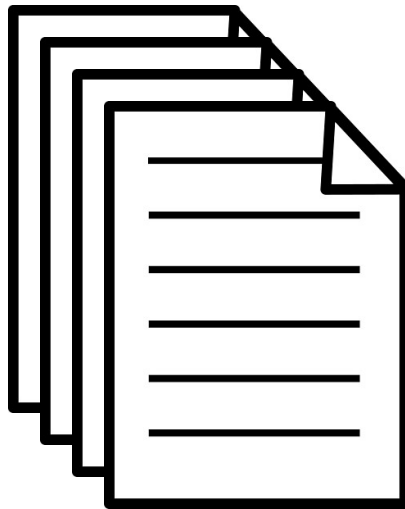


# Search-based Structured Prediction for biological event extraction

李辰

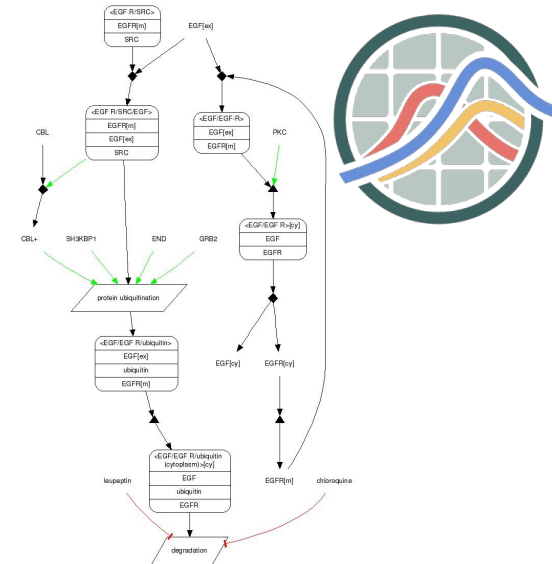
# Background

## Unstructured knowledge



## MEDLINE

## Structured knowledge



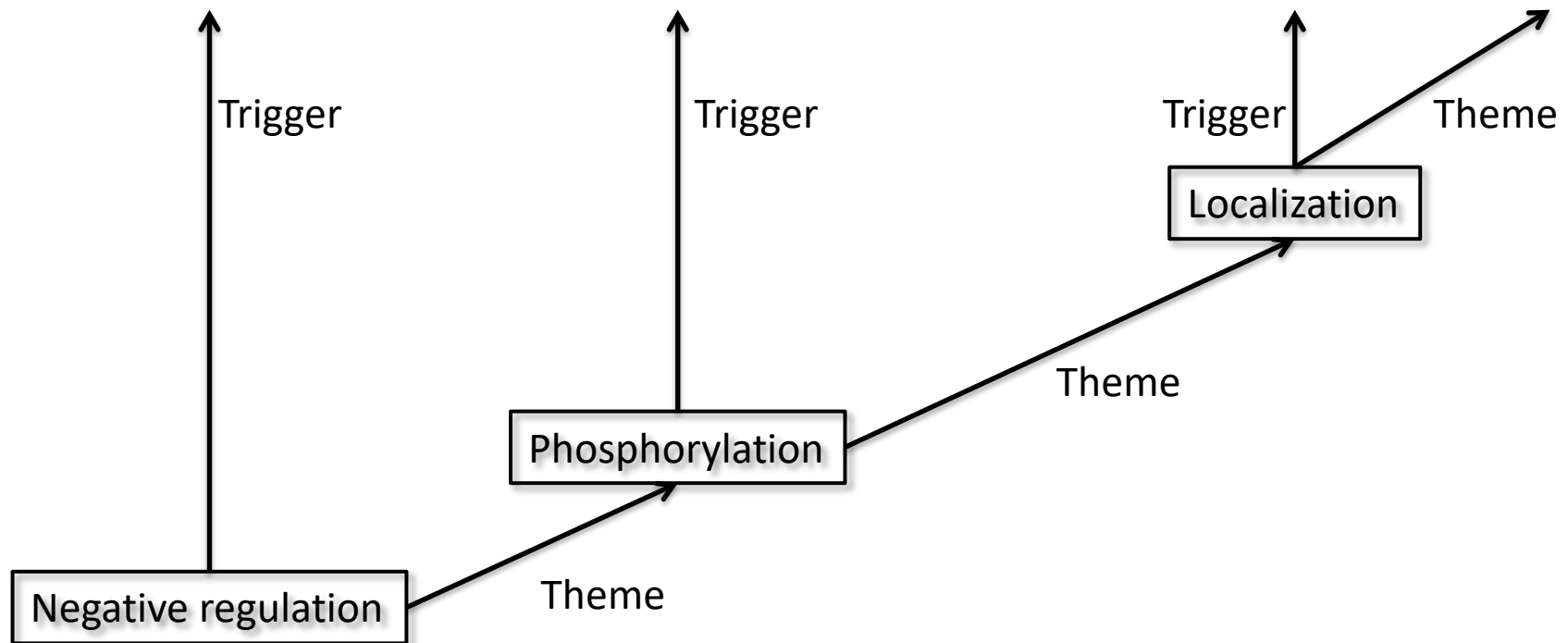
BioModels Database

Li C. et al., BMC systems biology

- Quantitatively characterise morphology of different types of biological networks in the scientific literature.
- Semantically enrich curated biological networks
- Discover hidden relations

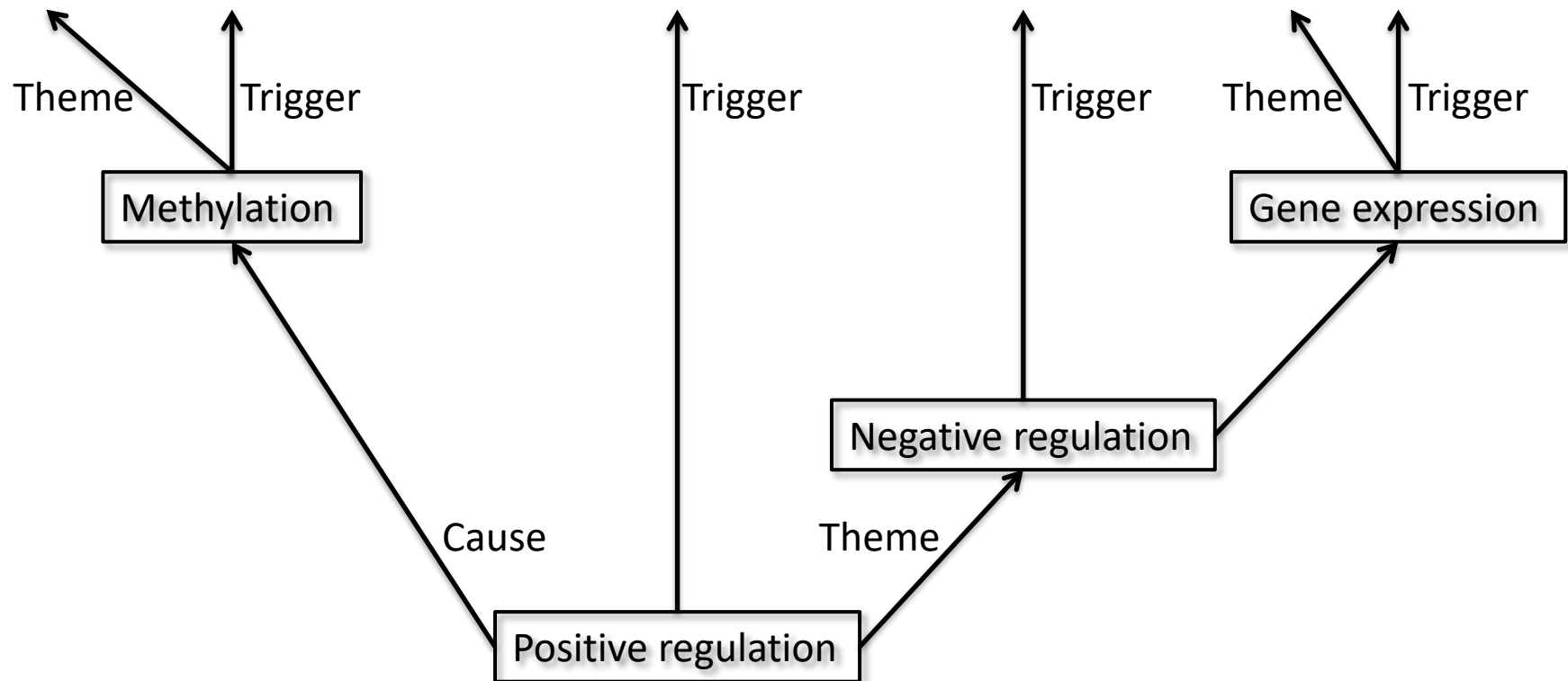
# Biomedical event extraction

decreased tyrosine phosphorylation and nuclear translocation of STAT6



# Biomedical event extraction

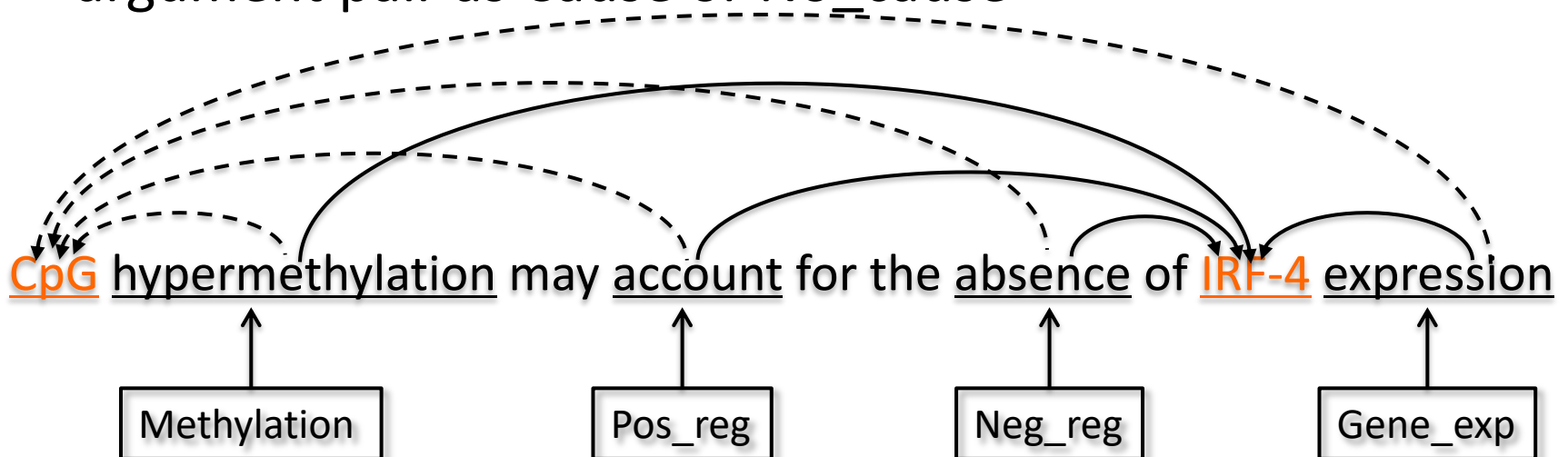
CpG hypermethylation may account for the absence of IRF-4 expression



# Modeling event extraction

## - Event extraction decomposition

- Trigger recognition: classify each token as one of the event types or No\_trigger
- Theme assignment: classify each candidate trigger-argument pair as Theme or No\_theme
- Cause assignment: classify each candidate trigger-argument pair as Cause or No\_cause



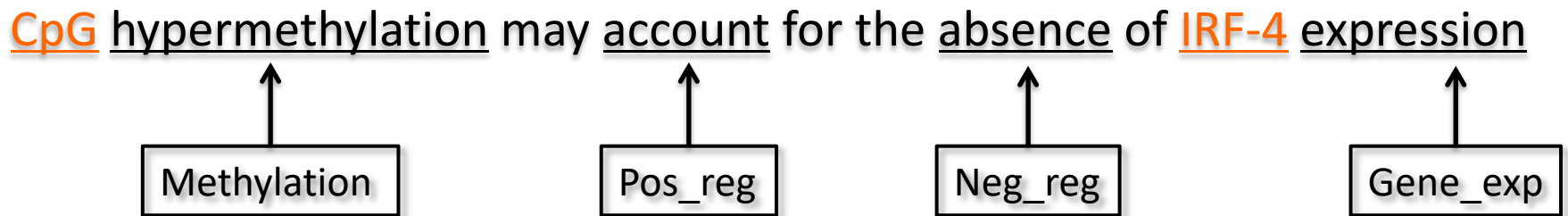
# Modeling event extraction

## - Event types and features

1. Simple events: an event takes one theme, e.g. gene expression
2. Binding: an event has multi-theme
3. Complex events: an event has both theme and cause
4. Recursive events: an event has theme and/or cause, which can be another event

# Modeling event extraction

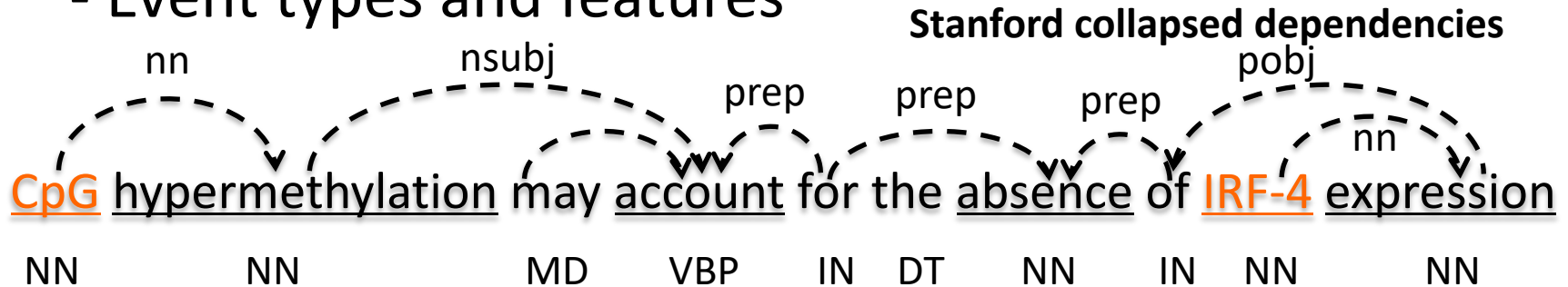
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## - Event types and features

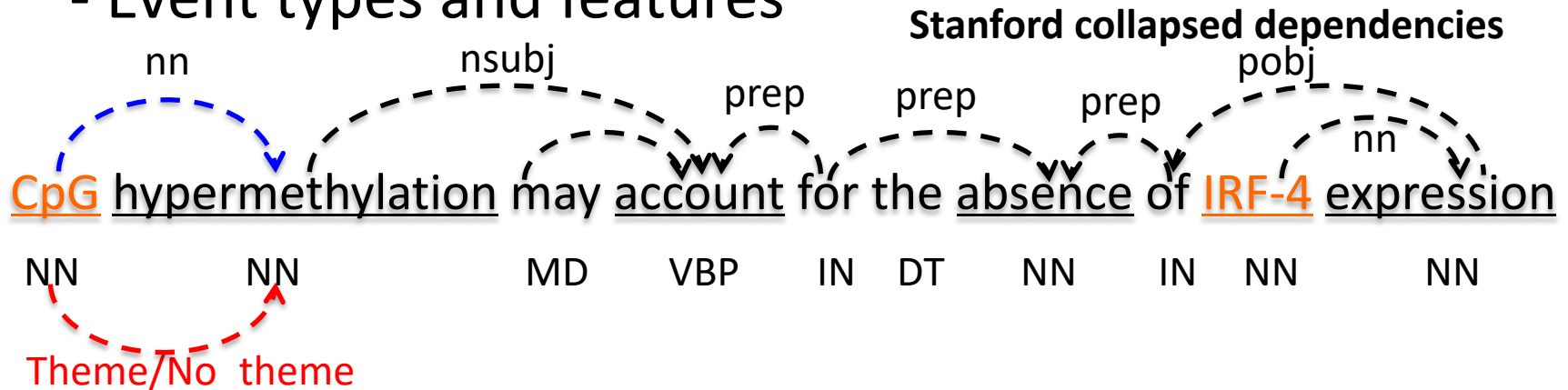


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# Modeling event extraction

## - Event types and features



**Path:** {trigger}<sup>nsubj</sup>→{argument}

**Lemma\_Path:** hypermethylation<sup>nsubj</sup>→{argument}

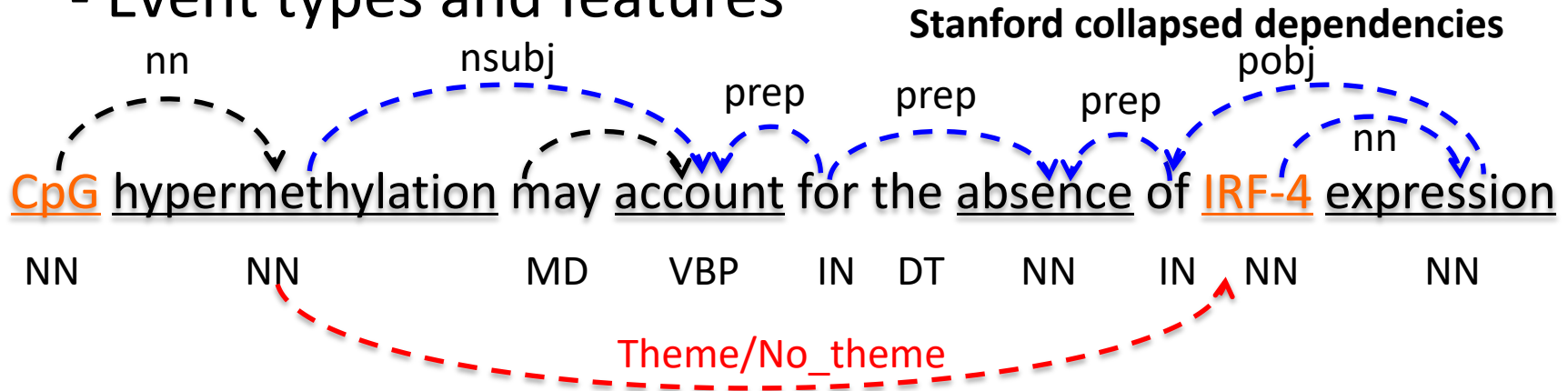
**POS\_Path:** NN<sup>nsubj</sup>→{argument}

*etc.*

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# Modeling event extraction

## - Event types and features



**Path:** {trigger}  $\xrightarrow{\text{nsubj}}$  {token}  $\xrightarrow{\text{prep}}$  {token}  $\xrightarrow{\text{prep}}$  {token}  $\xrightarrow{\text{prep}}$  {token}  $\xrightarrow{\text{pobj}}$  {token}  $\xrightarrow{\text{nn}}$  {argument}

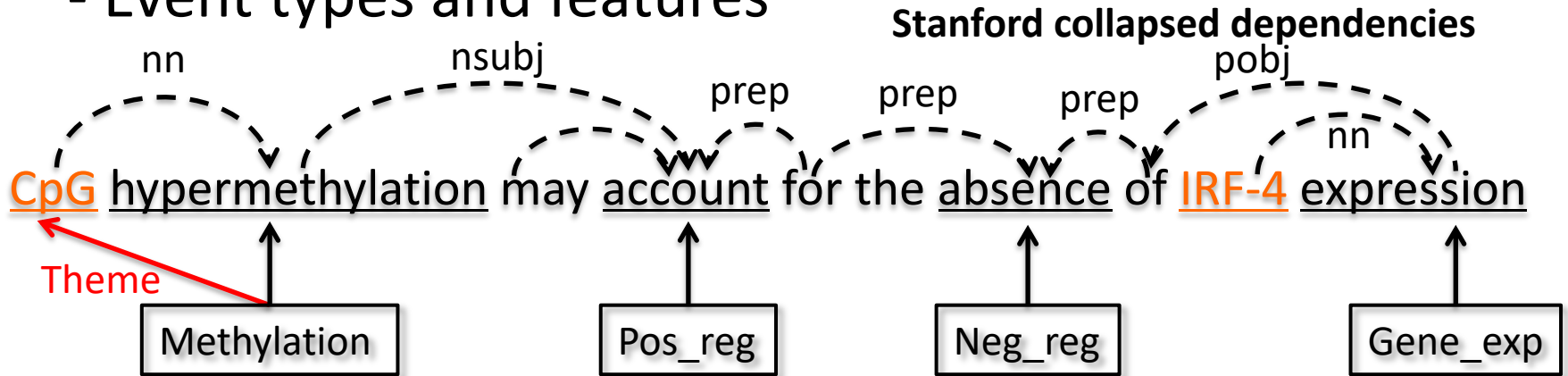
**Lemma\_Path:**  
hypermethylation  $\xrightarrow{\text{nsubj}}$  {token}  $\xrightarrow{\text{prep}}$  {token}  $\xrightarrow{\text{prep}}$  {token}  $\xrightarrow{\text{prep}}$  {token}  $\xrightarrow{\text{pobj}}$  {token}  $\xrightarrow{\text{nn}}$  {argument}

**POS\_Path:**  
hypermethylation  $\xrightarrow{\text{nsubj}}$  {token}  $\xrightarrow{\text{prep}}$  {token}  $\xrightarrow{\text{prep}}$  {token}  $\xrightarrow{\text{prep}}$  {token}  $\xrightarrow{\text{pobj}}$  {token}  $\xrightarrow{\text{nn}}$  {argument}  
etc.

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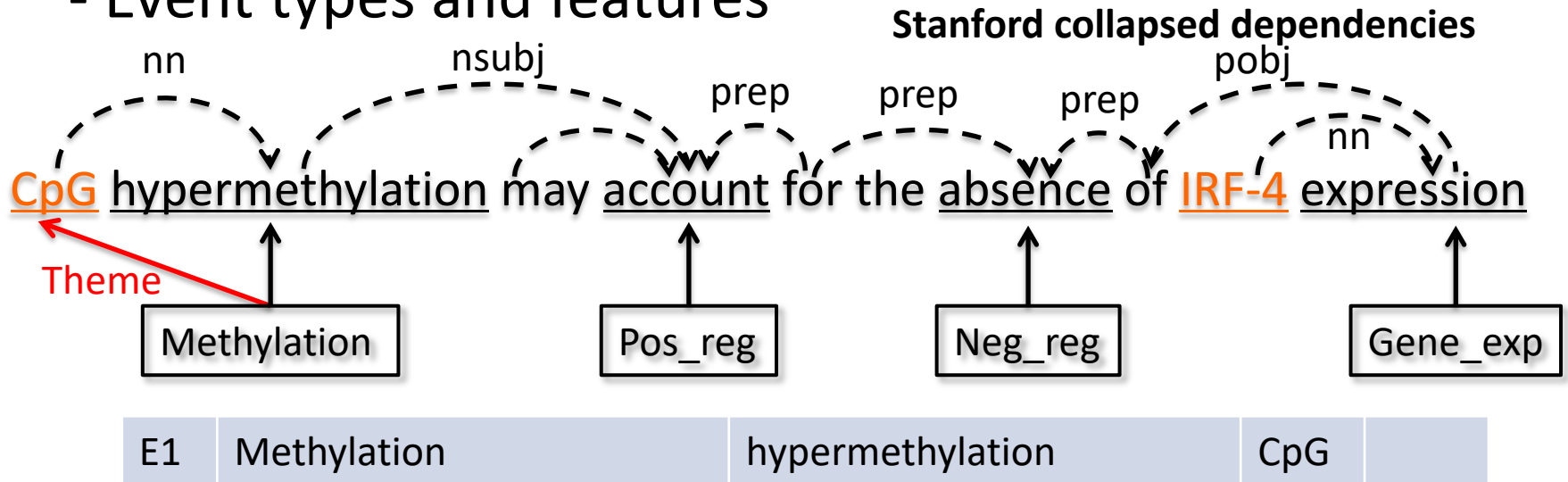
## - Event types and features



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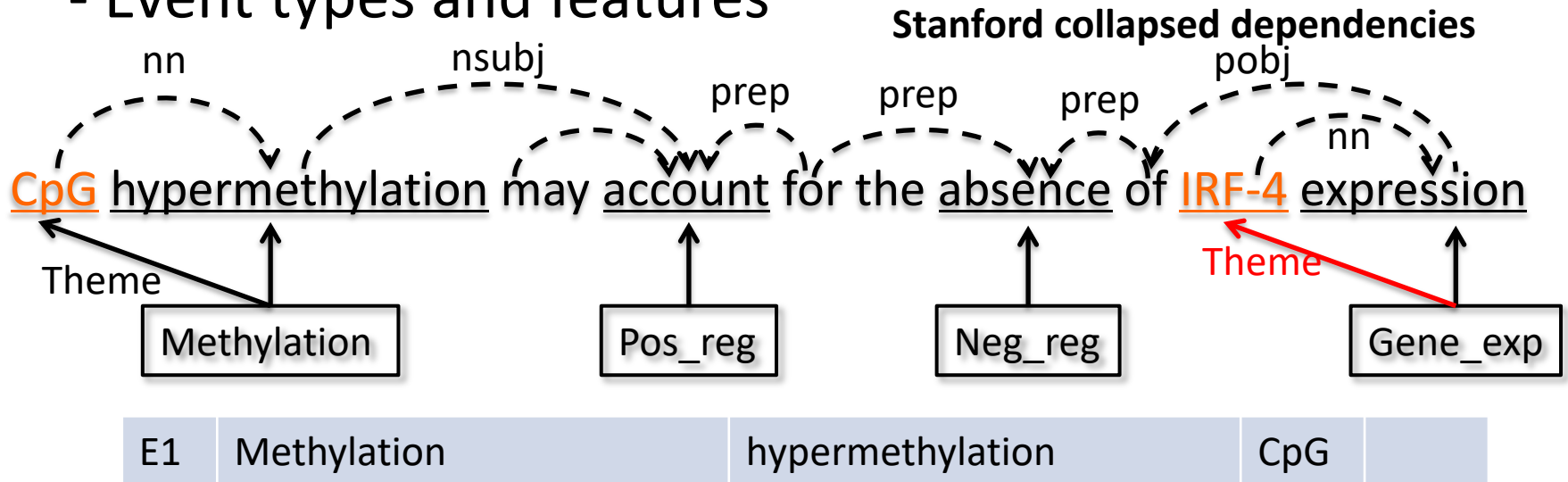
## - Event types and features



1. Simple events: an event takes one theme, e.g. gene expression
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# Modeling event extraction

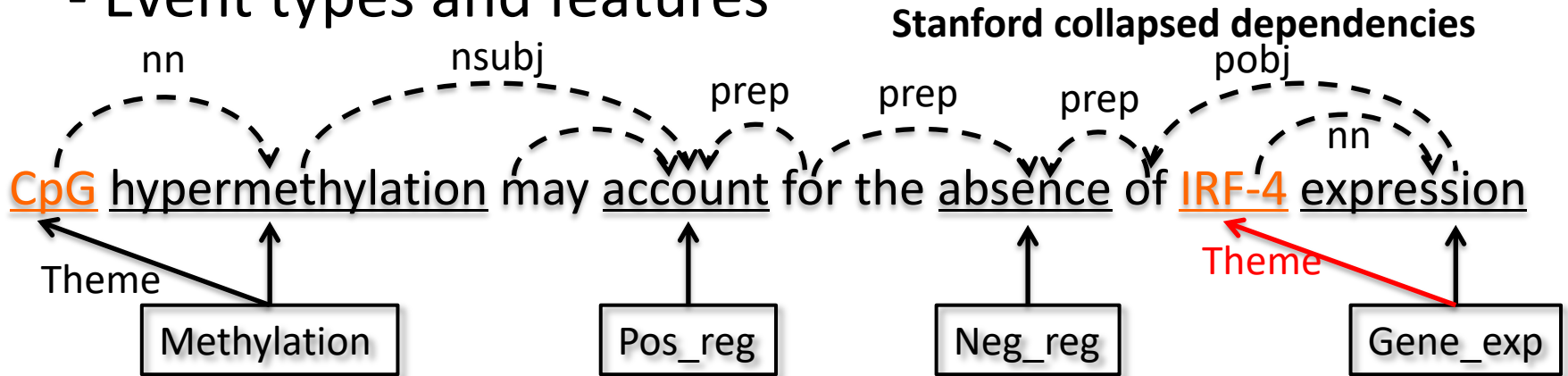
## - Event types and features



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# Modeling event extraction

## - Event types and features

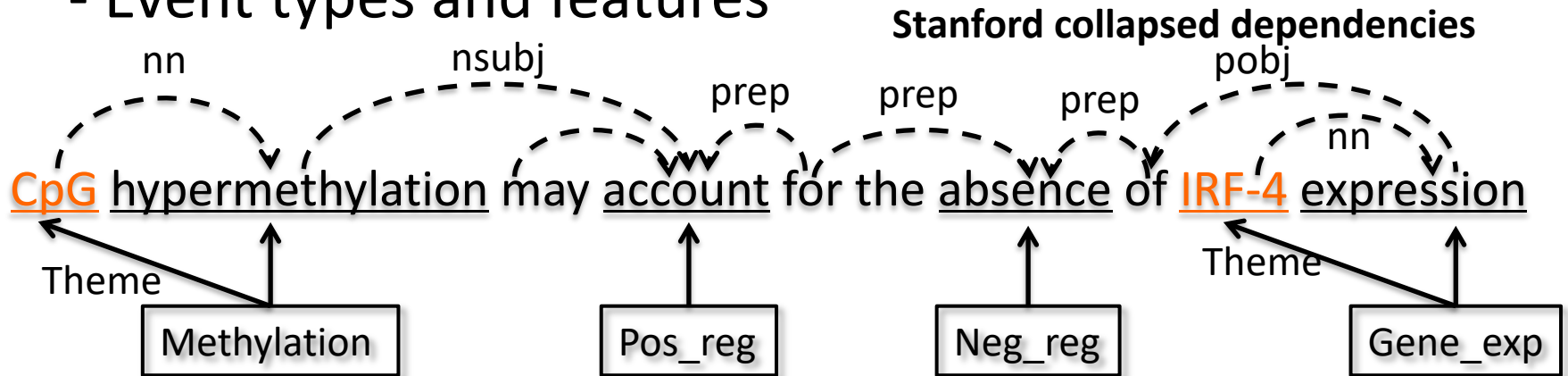


E1	Methylation	hypermethylation	CpG	
E2	Gene expression	expression	IRF-4	

1. Simple events: an event takes one theme, e.g. gene expression
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# Modeling event extraction

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# Modeling event extraction

- Event types and features

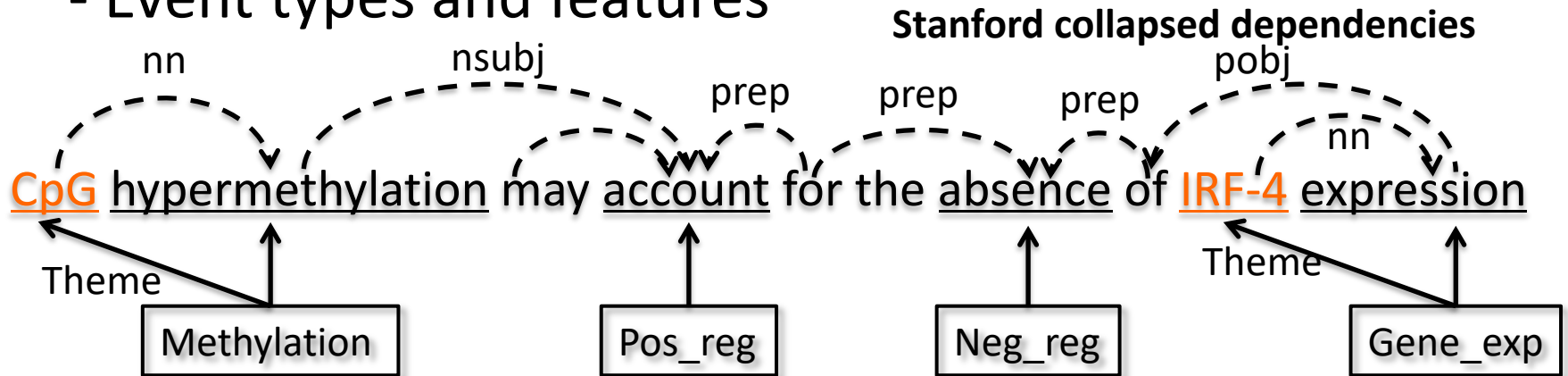
binding of A, B and C

1 <sup>st</sup> case	Binding	A
2 <sup>nd</sup> case	Binding	B
3 <sup>rd</sup> case	Binding	C
4 <sup>th</sup> case	Binding	AB
5 <sup>th</sup> case	Binding	BC
6 <sup>th</sup> case	Binding	AC
7 <sup>th</sup> case	Binding	ABC



# Modeling event extraction

## - Event types and features

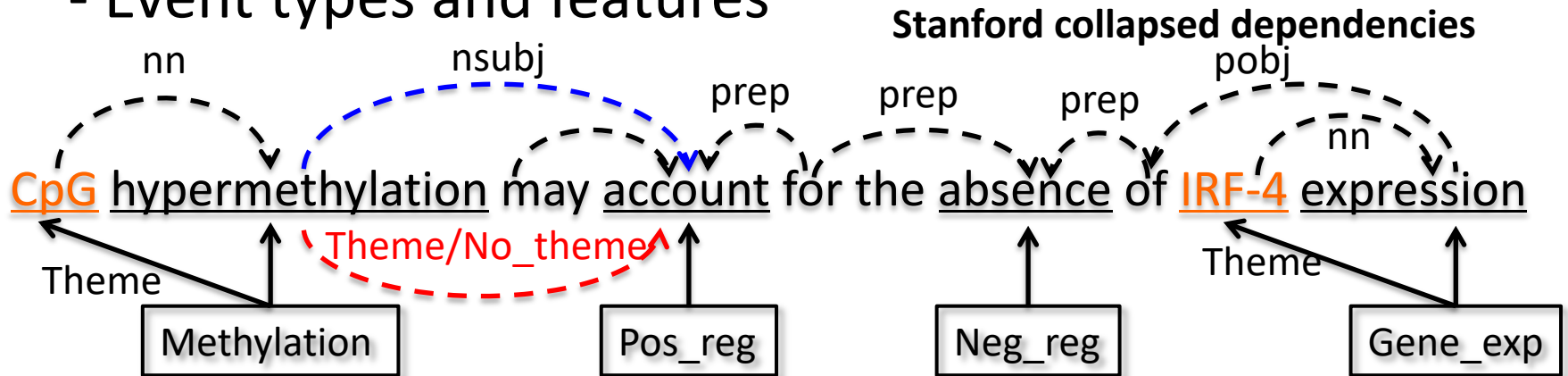


E1	Methylation	hypermethylation	CpG	
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# Modeling event extraction

## - Event types and features

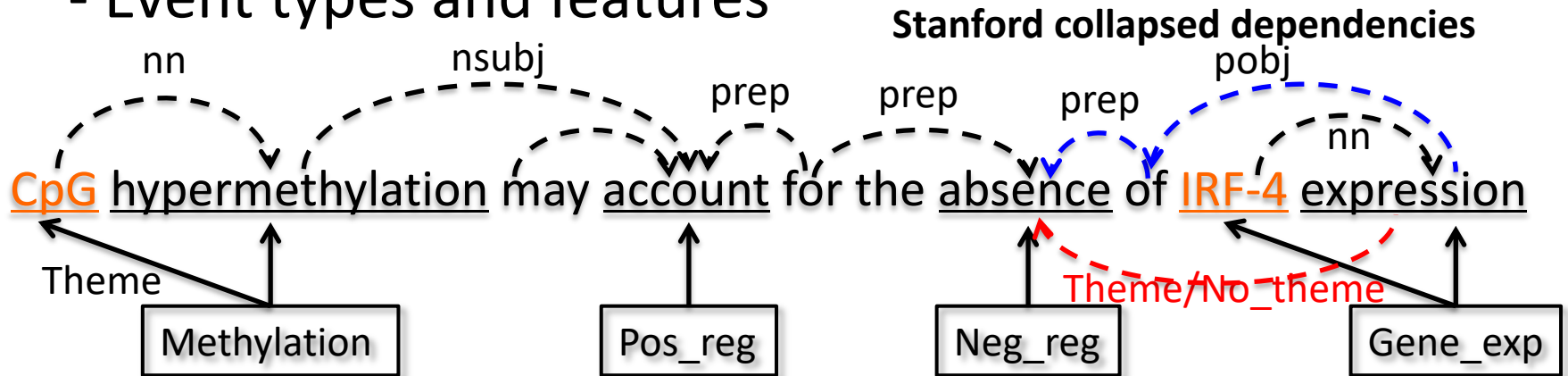


E1	Methylation	hypermethylation	CpG	
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# Modeling event extraction

## - Event types and features

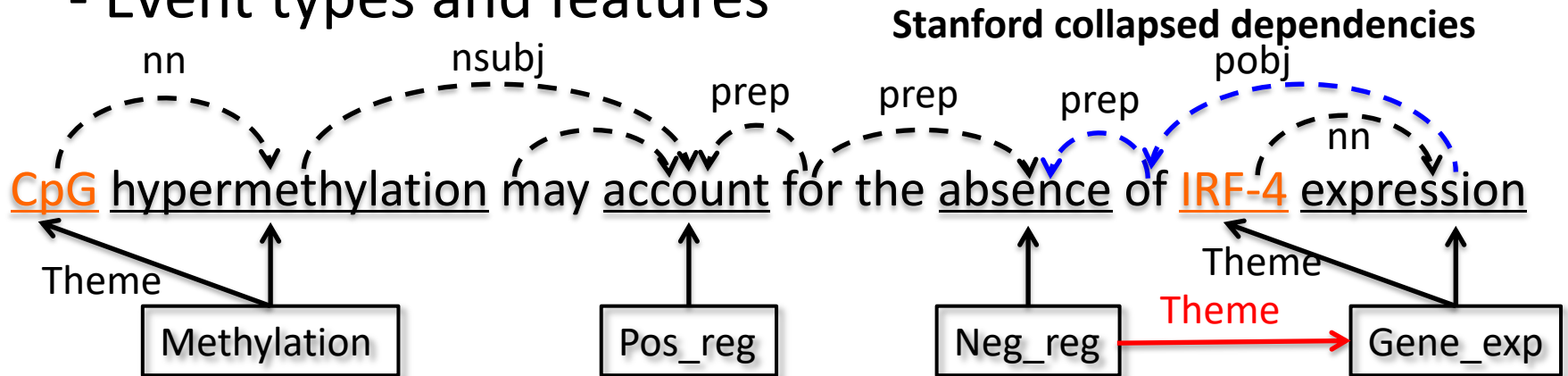


E1	Methylation	hypermethylation	CpG	
E2	Gene expression	expression	IRF-4	

1. Simple events: an event takes one theme, e.g. gene expression
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# Modeling event extraction

## - Event types and features

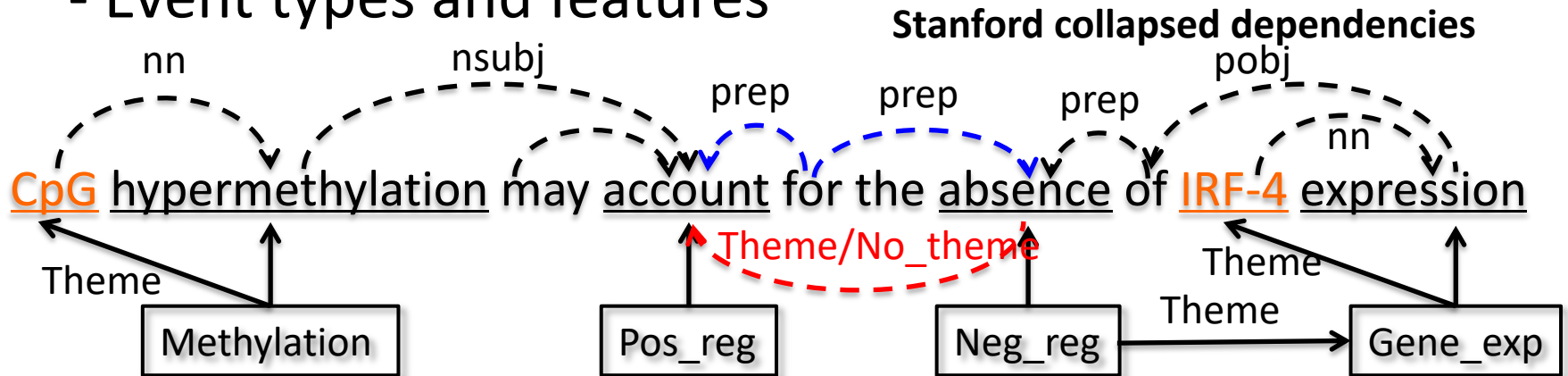


E1	Methylation	hypermethylation	CpG	
E2	Gene expression	expression	IRF-4	
E3	Negative regulation	absence	E2	

1. Simple events: an event takes one theme, e.g. gene expression
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# Modeling event extraction

## - Event types and features

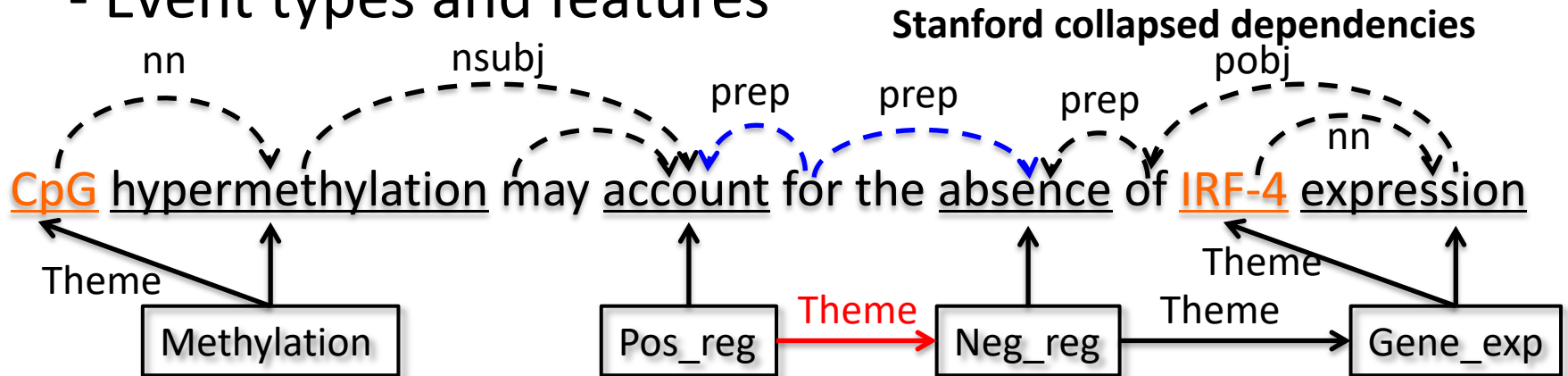


E1	Methylation	hypermethylation	CpG	
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# Modeling event extraction

## - Event types and features

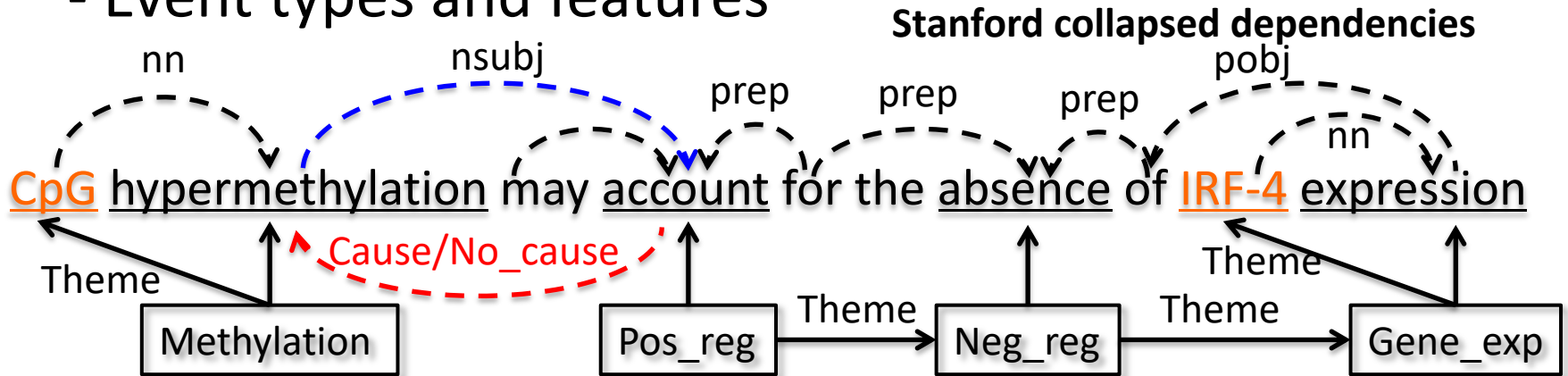


E1	Methylation	hypermethylation	CpG	
E2	Gene expression	expression	IRF-4	
E3	Negative regulation	absence	E2	
E4	Positive regulation	account	E3	

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# Modeling event extraction

## - Event types and features

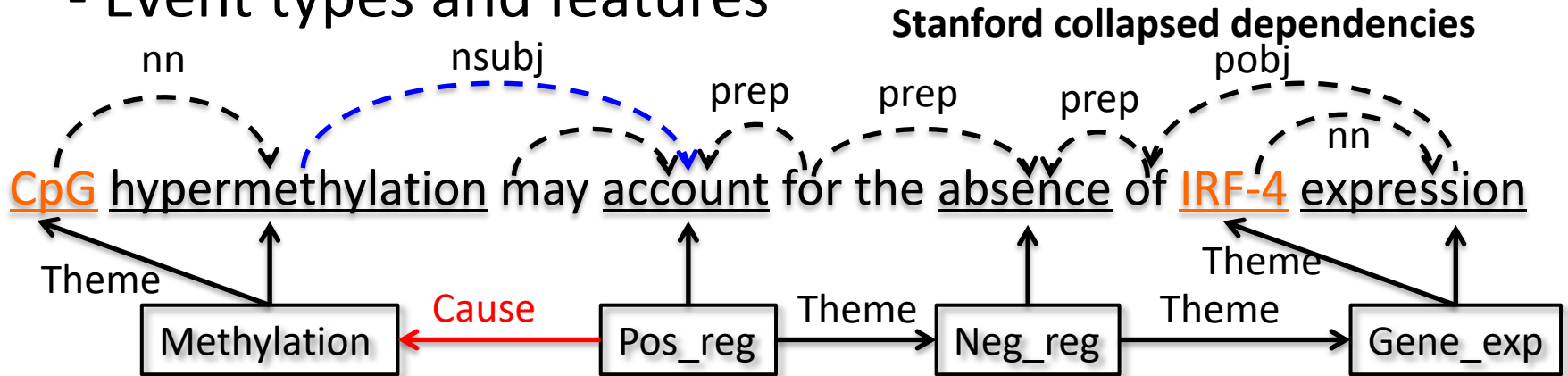


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# Modeling event extraction

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# BioNLP shared task

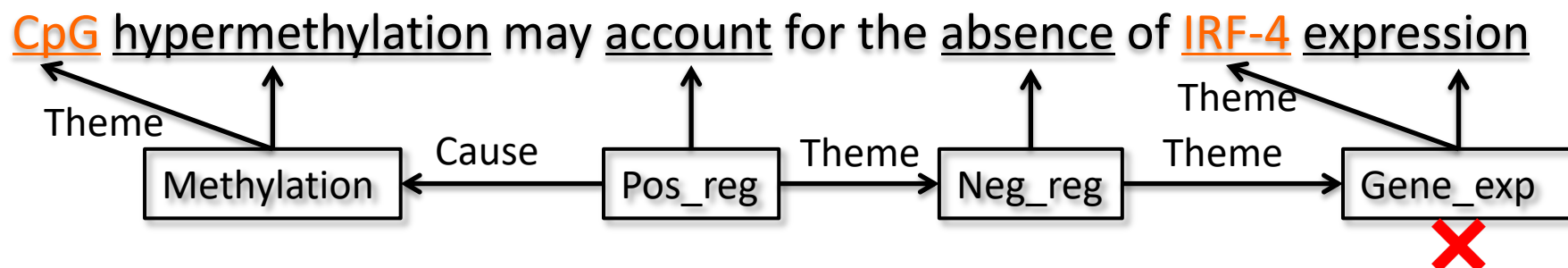
- Organized in 2009, 2011 and 2013 (next in 2015)
- Emphasis on
  - Expressive structured models of extracted information
  - “High-level” information extraction: directionality and polarity of reactions

## BioNLP shared task (*cont.*)

- Event structure is defined closer to reactions in biological networks.
  - Each event is associated with an event type
  - Event is explicitly mentioned in the text
  - Reactant number may vary in each event
  - Each reactant has different role, e.g. cause, theme, site etc.
  - Event can play a role in another event

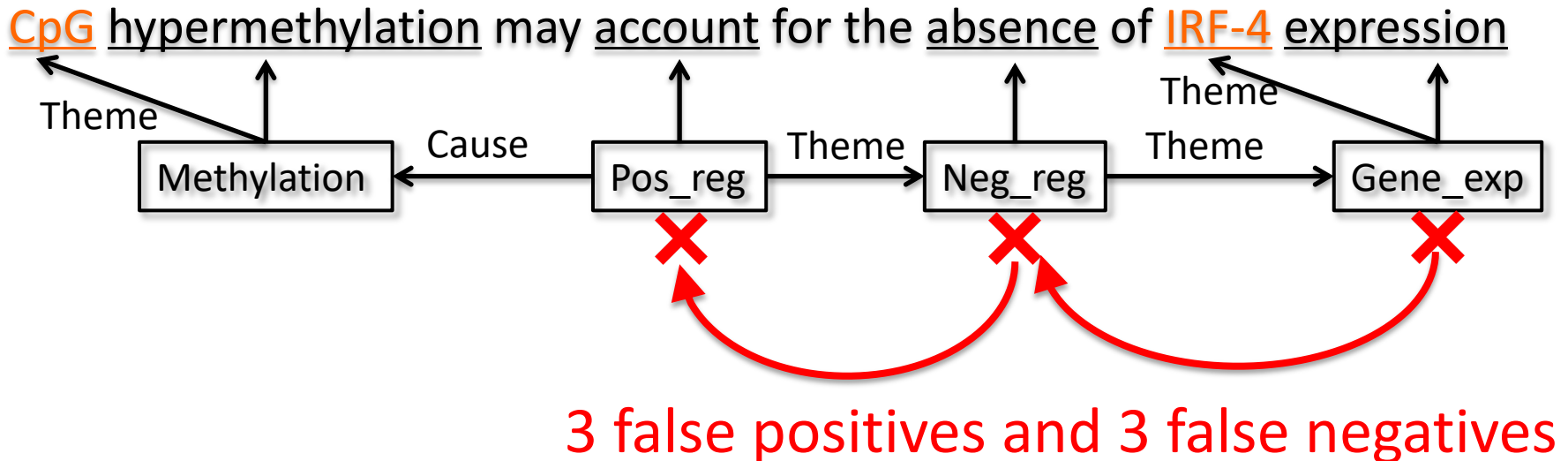
# BioNLP shared task (*cont.*)

- Structural evaluation



# BioNLP shared task (*cont.*)

- Structural evaluation



# Structured prediction requirements

- Labeled data
- Loss function
- Optimal policy
- A cost-sensitive classification

# Structured prediction (*algorithm*)

```
1 Initialise
2   Structured instances  $S$ ,
3   optimal policy  $\pi$ ,
4   cost sensitive learning algorithm  $CSCL$ 
5   loss function  $\ell$ 
6
7 Train
8   current policy  $h = \pi$ 
9   while  $h$  depends significantly on  $\pi$  do
10     Examples  $E = \emptyset$ 
11     for  $s$  in  $S$  do
12       Predict  $h(s) = \hat{y}_1 \dots \hat{y}_T$ 
13       for  $\hat{y}_t$  in  $h(s)$  do
14         Extract features  $\Phi_t = f(s, \hat{y}_{1:t-1})$ 
15         for each possible action  $y_t^i$  do
16           Predict  $y'_{t+1:T} = h(s | \hat{y}_{1:t-1}, y_t^i)$ 
17           Estimate  $c_t^i = \ell(\hat{y}_{1:t-1}, y_t^i, y'_{t+1:T})$ 
18         Add  $(\Phi_t, c_t)$  to  $E$ 
19     Learn a classifier  $h_{new} = CSCL(E)$ 
20      $h = \beta h_{new} + (1 + \beta)h$ 
21 Return policy  $h$ 
```

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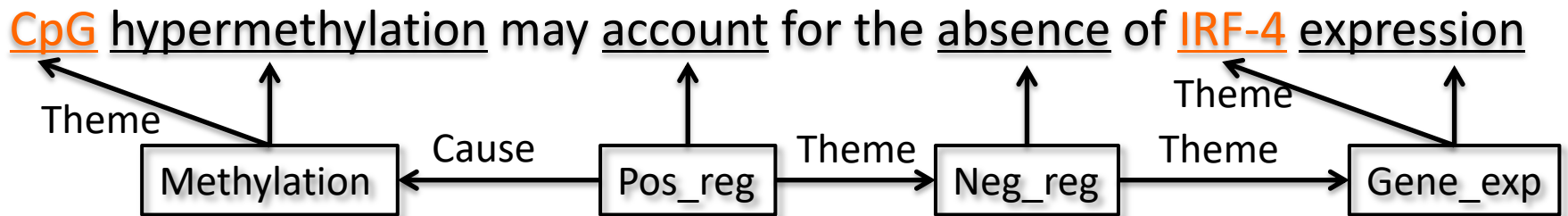
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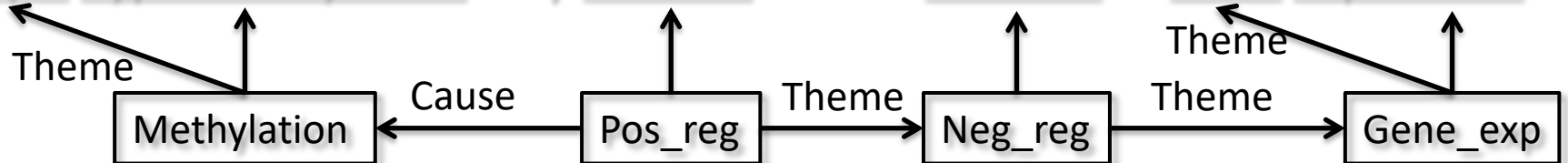
# Example of training



# Example of training

- 1<sup>st</sup> iteration -- Using the optimal policy

CpG hypermethylation may account for the absence of IRF-4 expression

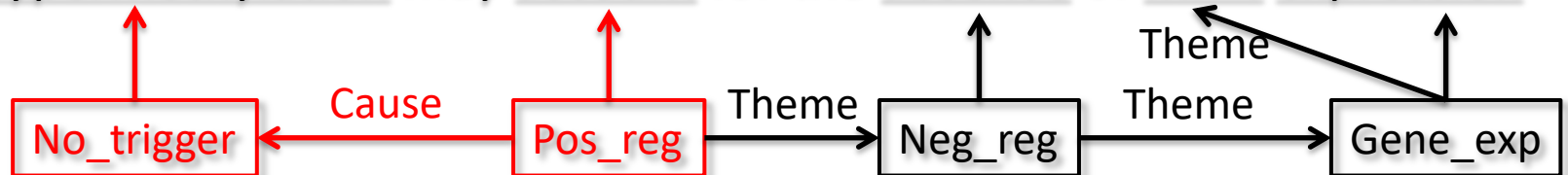


	Non_trigger	Pos_reg	Neg_reg	Methylation	Gene_exp
hypermethylation					
may					
account					
for					
absence					
expression					

# Example of training

- 1<sup>st</sup> iteration -- Using the optimal policy

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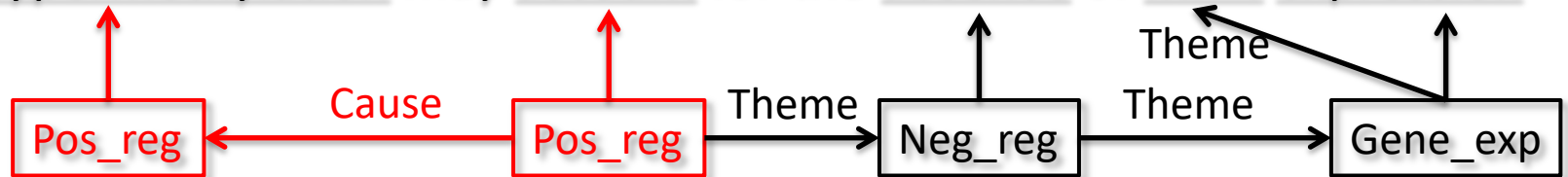


	Non_trigger	Pos_reg	Neg_reg	Methylation	Gene_exp
hypermethylation	3				
may					
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for					
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# Example of training

- 1<sup>st</sup> iteration -- Using the optimal policy

CpG hypermethylation may account for the absence of IRF-4 expression



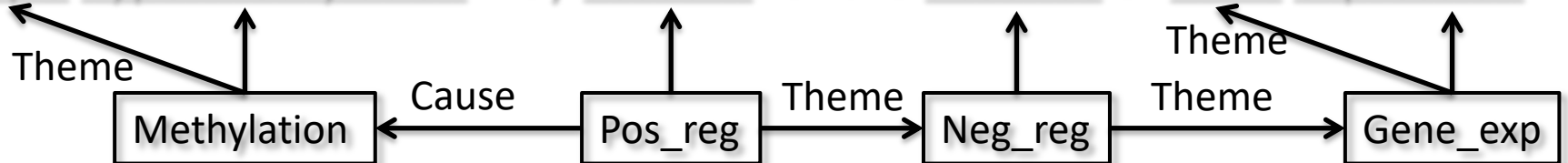
	Non_trigger	Pos_reg	Neg_reg	Methylation	Gene_exp
hypermethylation	3	4			
may					
account					
for					
absence					
expression					



# Example of training

- 1<sup>st</sup> iteration -- Using the optimal policy

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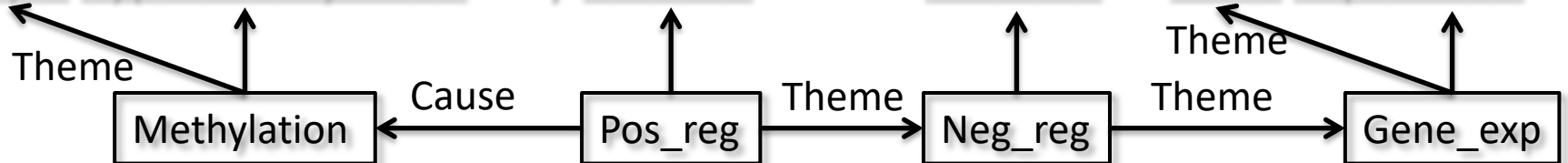


	Non_trigger	Pos_reg	Neg_reg	Methylation	Gene_exp
hypermethylation	3	4	4	0	4
may					
account					
for					
absence					
expression					

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- 1<sup>st</sup> iteration -- Using the optimal policy

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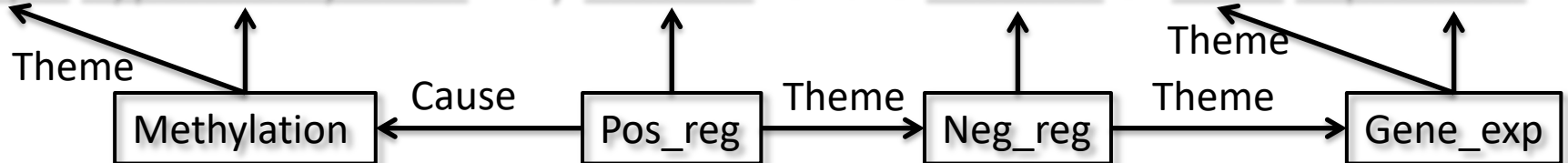


	Non_trigger	Pos_reg	Neg_reg	Methylation	Gene_exp
hypermethylation	3	4	4	0	4
may	0	0	0	0	0
account					
for					
absence					
expression					

# Example of training

- 1<sup>st</sup> iteration -- Using the optimal policy

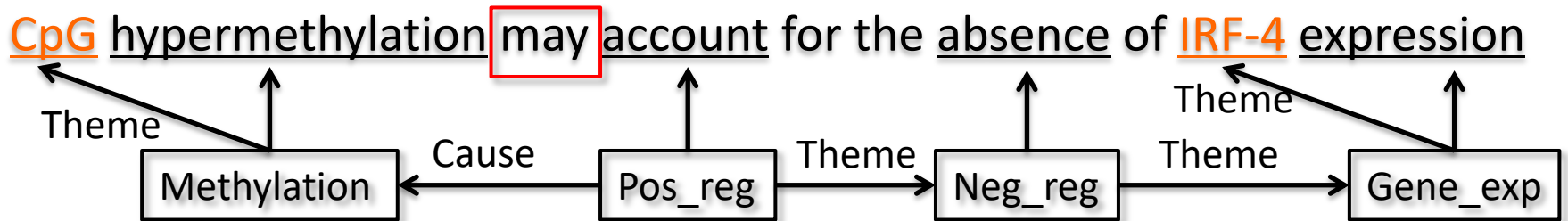
CpG hypermethylation may account for the absence of IRF-4 expression



	Non_trigger	Pos_reg	Neg_reg	Methylation	Gene_exp
hypermethylation	3	4	4	0	4
may	0	0	0	0	0
account	1	0	2	2	2
for	0	0	0	0	0
absence	3	4	0	4	4
expression	5	6	6	6	0

# Example of training

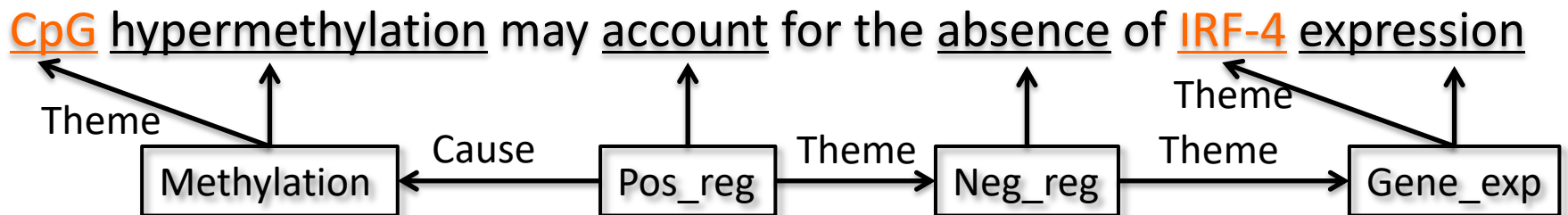
- 2<sup>nd</sup> iteration -- Learned hypothesis



	Non_trigger	Pos_reg	Neg_reg	Methylation	Gene_exp
hypermethylation	3	4	4	0	4
may	0	1	0	0	0
account					
for					
absence					
expression					

# Example of training

- 2<sup>nd</sup> iteration -- Learned hypothesis



	Non_trigger	Pos_reg	Neg_reg	Methylation	Gene_exp
hypermethylation	3	4	4	0	4
may	0	1	0	0	0
account	1	0	2	2	2
for	0	0	0	0	0
absence	3	4	0	4	4
expression	5	6	6	6	0

# Averaged perceptron

```
1 Input
2   training examples  $X = x_1 \dots x_T$ ,
3   cost vectors  $c_1 \dots c_T$ ,
4   initialise weights  $w_0^{(k)} = (0, \dots, 0)$ ,
5   for  $x_t \in X$  do
6     predict label  $\hat{y}_t = \operatorname{argmax}_k (w_t^{(k)} \times x_t)$ 
7     receive cost vector  $c_t$ 
8     if  $c_t^{(\hat{y}_t)} > 0$  then
9       loss  $\ell_t = w_t^{(\hat{y}_t)} \times x_t - w_t^{(y_t)} \times x_t + \sqrt{c_t^{(\hat{y}_t)}}$ 
10      learning rate  $\tau_t = \frac{\ell_t}{\|x_t\|^2 + \frac{1}{2c}}$ 
11      weight  $w_{t+1}^{(y_t)} = w_t + \tau_t x_t$ 
12              $w_{t+1}^{(\hat{y}_t)} = w_t - \tau_t x_t$ 
13  average  $w_{avg} = \frac{\sum_{i=0}^{T \times R} w_i}{T \times R}$ 
```

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```
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```

# Conclusion

- Biological networks are described in complicated structures in the scientific literature.
- Structured prediction for a complex structured prediction task
- Structured prediction achieved state-of-the-art performance
- A flexible system for extracting more types of biological networks



Questions?