

# A single-source k shortest paths algorithm to infer regulatory pathways in a gene network - Supplementary Material

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## List of Figures

- 1 The sorted e-values of the top GO term in  $GO_{ann}$  with different  $\lambda$ . . . . . 2

## List of Tables

- 1 The number of gold standards found by the  $k$  diverse paths algorithm with a significantly less e-value of a GO term in  $GO_{ann}$  . . . . . 2
- 2 The e-values of some enriched GO terms found by  $k = 5$  diverse short paths algorithm with different  $\lambda$ . . . . . 2
- 3 The top 10 genes and their corresponding paths found when  $\lambda = 0$  and 1.0. The starting node is RTS1. . . . . 3

(a) UNKNOWNCAUSAL using different  $\lambda$ .

(b) UNKNOWNTARGET using different  $\lambda$ .

Figure 1: The sorted e-values of the top GO term in  $GO_{ann}$  with different  $\lambda$ .  $k = 5$ .

Table 1: The number of gold standards found by the  $k$  diverse paths algorithm with a significantly less e-value of a GO term in  $GO_{ann}$  than the  $k$ -shortest paths algorithm ( $\lambda = 0$ ).  $k = 5$ .

$\lambda = 0.25$	<b>33</b>	$\lambda = 0.75$	<b>445</b>
$\lambda = 0.5$	<b>167</b>	$\lambda = 1.0$	<b>604</b>

Table 2: Case study - other examples: The e-values of some enriched GO terms found by  $k = 5$  diverse short paths algorithm with different  $\lambda$ . E-values larger than 0.01 are denoted by “-“, representing insignificance.

GO number	function name	$\lambda = 0$	$\lambda = 0.25$	$\lambda = 0.5$	$\lambda = 0.75$	$\lambda = 1.0$
Starting node: CMD1						
GO:0000780	condensed nuclear chromosome...	3.32E-3	2.61E-3	5.72E-4	2.43E-5	<b>8.51E-7</b>
*GO:0005934	cellular bud tip	-	-	-	-	<b>1.61E-4</b>
*GO:0006606	protein import into nucleus	5.28E-5	6.50E-5	1.55E-5	1.23E-8	-
GO:0010499	proteasomal ubiquitin-independent ...	1.42E-11	<b>1.39E-11</b>	4.05E-9	-	-
*GO:0031023	microtubule organizing center ...	<b>1.98E-9</b>	3.58E-9	2.55E-8	4.80E-8	3.20 E-8
*GO:0034504	protein localization in nucleus	-	-	1.02E-5	6.73E-9	<b>4.08E-12</b>
Starting node: RRP6						
GO:0000462	maturation of SSU-rRNA ...	3.00E-11	2.26E-11	5.42E-12	4.38E-4	-
*GO:0000466	maturation of 5.8S rRNA ...	<b>1.46E-31</b>	2.18E-31	1.67E-31	1.37E-21	6.20E-20
GO:0008094	DNA-dependent ATPase activity	-	-	-	3.53E-3	<b>4.92E-5</b>
*GO:0031126	snoRNA 3'-end processing	<b>1.10E-25</b>	2.27E-25	3.58E-25	1.76E-19	1.25E-20
Starting node: BUB3						
*GO:0000777	condensed chromosome kinetochore	2.53E-35	4.50E-35	<b>2.51E-36</b>	1.55E-3	5.03E-5
*GO:0007091	mitotic metaphase/anaphase transition	<b>9.36E-21</b>	1.67E-20	3.62E-18	7.30E-7	2.66E-11
GO:0016251	general RNA polymerase II transcription	-	-	-	<b>2.73E-9</b>	-
GO:0031145	anaphase-promoting complex-...	8.45E-13	<b>7.77E-13</b>	3.19E-11	-	4.19E-4
*GO:0033597	mitotic checkpoint complex	3.20E-4	8.94E-4	5.67E-3	1.05E-3	<b>6.11E-5</b>

\*These GO terms are in  $GO_{ann}$ , i.e., they annotate the starting node.

Table 3: The top 10 genes and their corresponding paths found when  $\lambda = 0$  and 1.0. The starting node is RTS1.

$\lambda = 0$			
1.PPH21	RTS1 → PPH21 RTS1 → RRD2 → PPH21 RTS1 → SGO1 → PPH21 RTS1 → RRD2 → TAP24 → PPH21 RTS1 → RRD2 → TPD3 → PPH21	6.PPH22	RTS1 → RRD2 → PPH22 RTS1 → SGO1 → PPH22 RTS1 → PPH21 → RRD2 → PPH22 RTS1 → RRD2 → TAP24 → PPH22 RTS1 → RRD2 → TPD3 → PPH22
2.RRD2	RTS1 → RRD2 RTS1 → PPH21 → RRD2 RTS1 → PPH21 → TAP24 → RRD2 RTS1 → SGO1 → PPH21 → RRD2 RTS1 → SGO1 → TPD3 → RRD2	7.SIT4	RTS1 → RRD2 → SIT4 RTS1 → PPH21 → RRD2 → SIT4 RTS1 → PPH21 → TOR2 → SIT4 RTS1 → RRD2 → TAP24 → SIT4 RTS1 → PPH21 → RRD1 → SIT4
3.SGO1	RTS1 → SGO1 RTS1 → PPH21 → SGO1 RTS1 → RRD2 → PPH21 → SGO1 RTS1 → RRD2 → TPD3 → SGO1 RTS1 → RRD2 → PPH22 → SGO1	8.RRD1	RTS1 → PPH21 → RRD1 RTS1 → RRD2 → TAP24 → RRD1 RTS1 → RRD2 → SIT4 → RRD1 RTS1 → RRD2 → PPH21 → RRD1 RTS1 → RRD2 → TPD3 → RRD1
4.TPD3	RTS1 → RRD2 → TPD3 RTS1 → SGO1 → TPD3 RTS1 → PPH21 → TPD3 RTS1 → PPH21 → RRD2 → TPD3 RTS1 → PPH21 → RRD1 → TPD3	9.TOR2	RTS1 → PPH21 → TOR2 RTS1 → RRD2 → TAP24 → TOR2 RTS1 → RRD2 → SIT4 → TOR2 RTS1 → RRD2 → PPH21 → TOR2 RTS1 → PPH21 → TAP24 → TOR2
5.TAP24	RTS1 → RRD2 → TAP24 RTS1 → PPH21 → TAP24 RTS1 → PPH21 → RRD2 → TAP24 RTS1 → PPH21 → TOR2 → TAP24 RTS1 → RRD2 → SIT4 → TAP24	10.CDC55	RTS1 → PPH21 → CDC55 RTS1 → RRD2 → PPH21 → CDC55 RTS1 → RRD2 → TPD3 → CDC55 RTS1 → RRD2 → PPH22 → CDC55 RTS1 → PPH21 → ZDS2 → CDC55
$\lambda = 1.0$			
1.PPH21	RTS1 → PPH21 RTS1 → RRD2 → PPH21 RTS1 → SGO1 → PPH21	6.TAP24	RTS1 → RRD2 → TAP24 RTS1 → PPH21 → TAP24
2.RRD2	RTS1 → RRD2 RTS1 → PPH21 → RRD2 RTS1 → SGO1 → TPD3 → RRD2	7.ZDS2	RTS1 → PPH21 → ZDS2 RTS1 → RRD2 → TPD3 → ZDS2 RTS1 → SGO1 → TPD3 → CDC55 → ZDS2
3.SGO1	RTS1 → SGO1 RTS1 → PPH21 → SGO1 RTS1 → RRD2 → TPD3 → SGO1	8.PPH22	RTS1 → RRD2 → PPH22 RTS1 → SGO1 → PPH22
4.TPD3	RTS1 → RRD2 → TPD3 RTS1 → SGO1 → TPD3 RTS1 → PPH21 → TPD3	9.SIT4	RTS1 → RRD2 → SIT4 RTS1 → PPH21 → TOR2 → SIT4 RTS1 → PPH21 → TOR2 → SIT4
5.CDC55	RTS1 → PPH21 → CDC55 RTS1 → RRD2 → TPD3 → CDC55 RTS1 → SGO1 → PPH22 → CDC55	10.TOR2	RTS1 → PPH21 → TOR2 RTS1 → RRD2 → TAP24 → TOR2