**Supplementary Figures to** 

## Steady state statistical correlations predict bistability in reaction motifs

Suchana Chakravarty and Debashis Barik\*

School of Chemistry, University of Hyderabad, Central University P.O., Hyderabad, 500046, Telangana, India

\*Corresponding author, Email address: <u>dbariksc@uohyd.ac.in</u>



**Supplementary Figure 1:** Bimodality of steady state distributions, stochastic analogue of deterministic bifurcation diagram. The steady state distributions of X are plotted for a range of values of the synthesis rate ( $k_{s,i}$ ) from stochastic simulations for the models with indicated strengths of positive feedback. Top row: *DNFL-s*, middle row: *PFL-s* and bottom row: *PFL-d*. The corresponding deterministic bifurcation diagrams are overlaid on top of the distribution.



**Supplementary Figure 2:** 3<sup>rd</sup> order vs 2<sup>nd</sup> order cumulants for four different models without intrinsic bistability.



**Supplementary Figure 3:** a) Schematic diagram of gene expression model. The rate constant of each elementary reaction is mentioned in the diagram. b) Variance of protein (P) is correlated with the covariance between protein and mRNA (M) for varied range of translation rate constant ( $k_p$ ) or degradation rate constant ( $\gamma_p$ ). While the translational rate is varied the variance-covariance shows is cusp-shaped structure (red line) falsely indicating bistability. However, when degradation rate constant of protein is varied the cusp-shape disappears confirming monostability. (d) Similar conclusion are made based on plot of 4<sup>th</sup> order cross-cumulant with 3<sup>rd</sup> order cross-cumulant.



**Supplementary Figure 4:** Various statistical analysis of *DNFL-d* model with added Gaussian noise (zero mean). The strengths of the added noise are indicated inside the plot. The positive feedback strength is chosen to be  $k_{a,1} = 0.01$ .



**Supplementary Figure 5:** Various statistical analysis of *PFL-s* model with added Gaussian noise (zero mean). The strengths of the added noise are indicated inside the plot. The positive feedback strength is chosen to be  $k_{a,3} = 0.16$ .