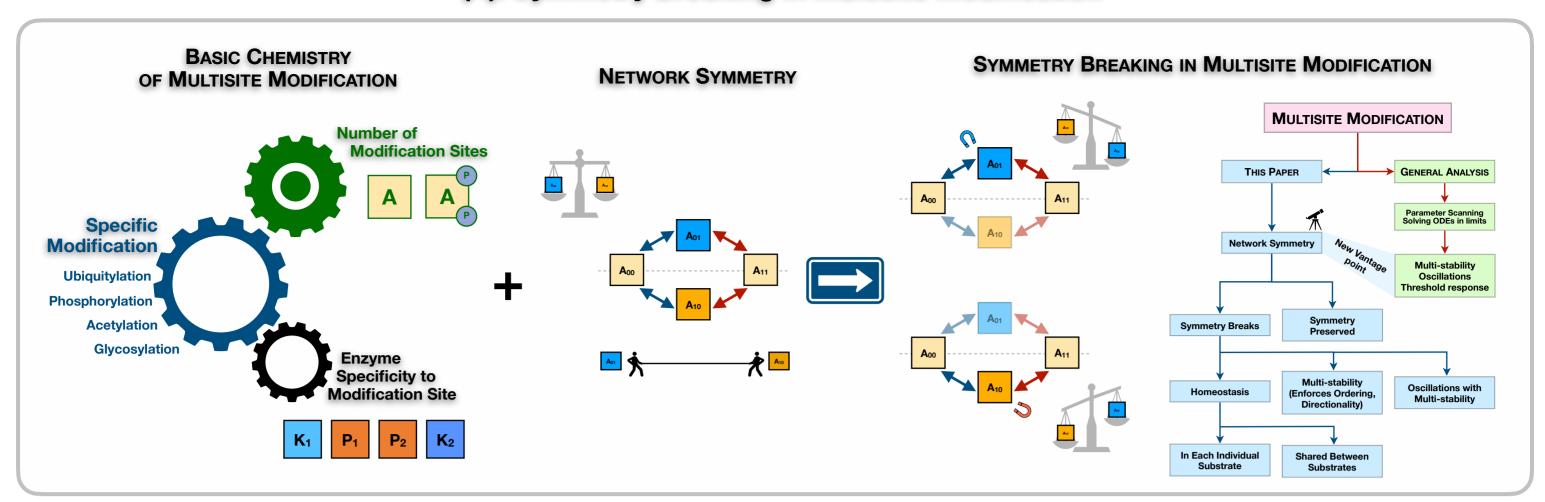
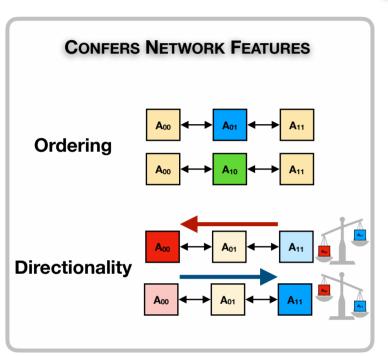
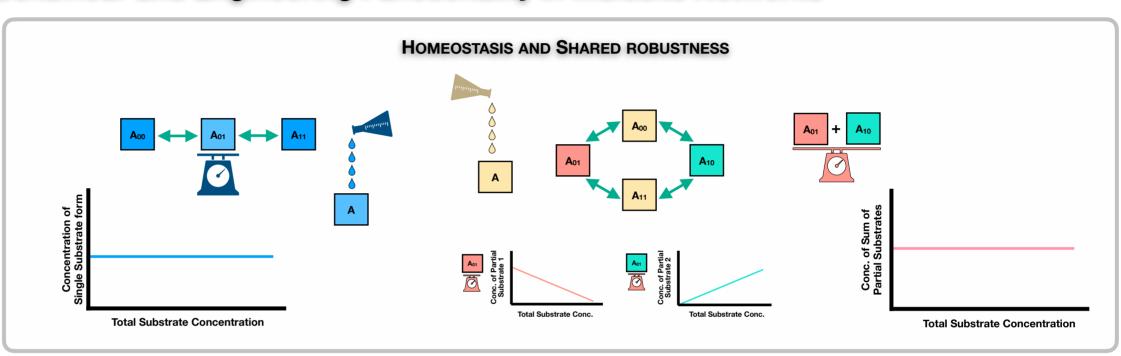
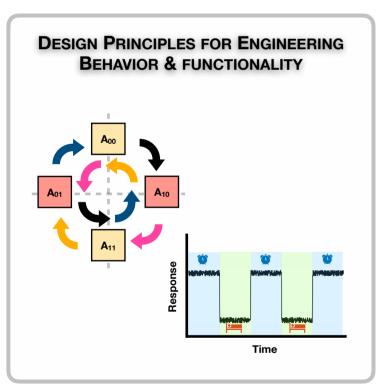
(A). Symmetry Breaking in Multisite Modification



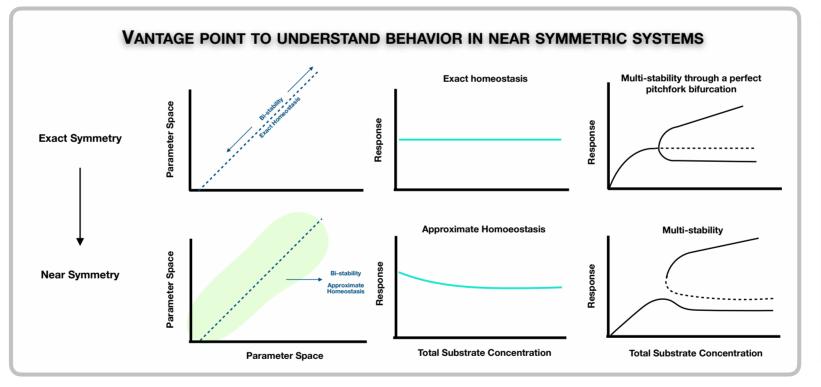
(B). Symmetry & Symmetry Breaking as a Focal Point for Understanding Behaviour and Engineering Functionality in Multisite Networks

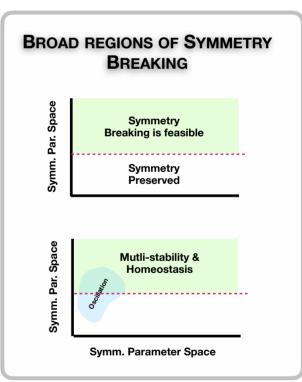






Dephosphorylation)





	(C). Summary of Symmetry and Symmetry Breaking Behavior in MSP Networks						
		Case 1		Case 2		Case 3	
Ordered DSP (Common Kinase Common Phosphatase)	Yes	Partial substrate form (A _p) exhibits concentration robustness post symmetry breaking					
Random System 1 (Common Kinase Common Phosphatase)	Yes	Partial substrate forms (A ₀₁ & A ₁₀) individually exhibit concentration robustness post symmetry breaking		No	Yes	Sum of the partially modified substrates (A ₀₁ + A ₁₀) exhibit concentration robustness post symmetry breaking	
Random System 2 (Separate Kinase Common Phosphatase)			No				
Random System 3 (Separate Kinase Separate Phosphatase)	Yes	Partial substrate forms (A ₀₁ & A ₁₀) individually exhibit concentration robustness post symmetry breaking	Yes	Completely modified and the unmodified substrates (A ₀₀ & A ₁₁) individually exhibit concentration robustness post symmetry breaking	Yes*	No exact robustness exhibited (Approximate concentration robustness in pairs of substrate possible - see text)	
Mixed Random 1 (Common Kinase Common Phosphatase Distributive Phosphorylation Processive Dephosphorylation)				No			
Mixed Random 2 (Separate Kinase Common Phosphatase Distributive Phosphorylation Processive			Yes	Completely modified and the unmodified substrates (A ₀₀ & A ₁₁) individually exhibit concentration robustness post symmetry breaking			