

# Systems Biology Research Symposium

## Oral Presentation Session

Grand Ballroom  
Tuesday, June 5th  
7:00-8:30pm

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Dissecting Hierarchical Regulatory Network of Estrogen-Dependent Breast Cancer through an Integrative Genomic Analysis

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A global level profiling of *in vivo* protein-DNA interactions using ChIP-based technologies has evolved rapidly in recent years, the emerging technology ChIP-seq has shown many advantages in terms of resolutions, accuracy, and cost. In this study, we have combined ChIP-seq and computational approaches to dissect regulatory networks for estrogen-dependent breast cancer. Although many studies including ours have genome-widely identified thousands of ER $\alpha$  binding sites and revealed a few transcription factor (TF) partners with ER $\alpha$ , such as AP1, FOXA1, MYC, SP1, little has been explored about complex regulatory networks. Our integrated genomic analysis reveals hierarchical regulatory networks existing for MCF7 and tamoxifen-resistant MCF7 (MCF7-T) cells. The fact that only 20 common genes showing differential expression after E2-induced and few common TF partners between MCF7 and MCF7-T strongly suggests they underlie completely different estrogen-mediated mechanisms. Our results not only confirmed estrogen-mediated gene repression path requiring ER $\alpha$  binding sites concluded from other studies, but also revealed a new estrogen-mediated up-regulated path requiring XBP1. Our data provide additional resources of ER $\alpha$  binding sites and regulated genes, thus may guide further study for underlying mechanisms in breast cancer.