A Probablistic Model for Characterizing Regulatory Targets of Transcription Factors from ChIP-Seq and ChIP-chip Binding Profiles

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To be submitted to Bioinformatics very soon

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• Suppose that we have a set of genes $\{g\}$, a set of Transcription Factors (TF) $\{t\}$, and the binding profile of each TF t on each gene g, that is, S_g^t , we want to calculate the posterior probability of TF t's targeting gene g,

$$p(t_{g} = 1 \mid S_{g}^{t}) = \frac{p(t_{g} = 1, S_{g}^{t})}{p(S_{g}^{t})} \propto p(t_{g} = 1, S_{g}^{t}) \approx \sum_{i=1}^{i=n} p(t_{gi} = 1, S_{gi}^{t})$$

$$= \sum_{i=1}^{i=n} p(t_{gi} = 1) p(S_{gi}^{t} \mid t_{gi} = 1)$$

$$\propto \sum_{i=1}^{i=n} w_{gi} \times f(S_{gi}) = \sum_{i=1}^{i=n} w_{i} \times f(S_{gi})$$

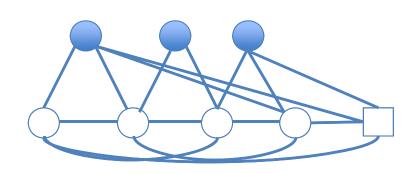
 \mathcal{W}_{gi} is the prior prob. that TF \boldsymbol{t} targets position \boldsymbol{i} of gene \boldsymbol{g} , and we assume that all the genes share the same position-specific prior probabilities w_i , which is estimated by Chao by the proportion of the total amount of reads or peak heights covered at position \boldsymbol{i} over all genes

Discriminative Semi-RBMs for Modeling Interactions between Histone Modifications and TF Bindings to Predict Gene Expression

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Semi-RBMs with Lateral Connections





visible units

label

$$E(v,h) = -\sum_{i,j} v_{i}h_{j}W_{ij}$$

$$-\sum_{y,j} l_{y}h_{j}W_{yj} - \sum_{y} l_{y}v_{i}v_{i}L_{ii} - \sum_{i} v_{i}b_{i} - \sum_{j} h_{j}c_{j} - \sum_{y} l_{y}d_{y}$$

$$\langle v_i h_j \rangle^0$$
 $\langle v_i h_j \rangle^1$
 $t = 0$ $t = 1$ reconstruction

$$p(v,h) \propto e^{-E(v,h)}$$

$$\Delta w_{ij} = \varepsilon \left(\langle v_i h_j \rangle^0 - \langle v_i h_j \rangle^1 \right)$$

Take home message: given expression level, the model predicts that, a small number of highly interacted histone mark features are essential, and all the others are redundant by-products.