
Appendix: Cross Species Expression Analysis using a Dirichlet Process Mixture Model with Latent Matchings

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1 Posterior distribution of μ_Y, Λ_Y :

$$\begin{aligned} \kappa_{Yk} &= \kappa_{Y0} + n_Y & m_Y &= \frac{1}{\kappa_Y}(\kappa_{Y0}m_{Y0} + n_Y\bar{y}) \\ S_Y^{-1} &= S_{Y0}^{-1} + V_Y + \frac{\kappa_{Y0}n_Y}{\kappa_{Y0} + n_Y}(\bar{y} - m_{Y0})(\bar{y} - m_{Y0})^T & \nu_Y &= \nu_{Y0} + n_Y \end{aligned}$$

where $n_Y = \sum_{i=1}^{n_x} \sum_{j=1}^{n_y} \psi_{i,j,k}$, $\bar{y} = \frac{1}{n_Y} \sum_{i=1}^{n_x} \sum_{j=1}^{n_y} \psi_{i,j,k} y_j$ and $V_Y = \sum_{i=1}^{n_x} \sum_{j=1}^{n_y} \psi_{i,j,k} (y_j - \bar{y})(y_j - \bar{y})^T$.

2 Data Normalization

As noted in the paper, the biological data was obtained from two previous publications. In both cases the authors have deposited normalized expression values in public databases (Stanford Microarray Database and Gene Expression Omnibus) and we have retrieved these datasets from these public databases. The data was indeed log2 transformed to obtain log ratios for changes vs. control (uninfected cells in both cases). These are standard procedures in microarray analysis and we would include this description in the updated paper. Parameters for BLASTN are based on the NCBI recommendations.

3 GO enrichment analysis for clusters inferred by DPMMLM

Table 1, 2, 3 and 4 present the GO enrichment analysis for cluster 2, 3, 4, 5 inferred by DPMMLM.

4 GO enrichment analysis for clusters inferred by DPMM

To test whether these differences inferred by the algorithm are biologically meaningful we compared our Dirichlet method to a method that uses deterministic assignments, as was done in the past (DPMM). Using such assignments the algorithm identified only two clusters. Neither of these clusters looked homogenous across species. Table 5 and 6 show the GO enrichment result for the two identified clusters.

P value	Corrected P val	GO term description	P value	Corrected P val	GO term description
2.98216e-18	<0.001	regulation of transcription, DNA-dependent	9.80701e-11	<0.001	protein binding
1.58764e-16	<0.001	ion binding	1.71813e-10	<0.001	hydrogen ion transmembrane transporter activity
1.28856e-16	<0.001	cation binding	8.76251e-10	<0.001	energy coupled proton transport, down electrochemical gradient
4.81459e-16	<0.001	zinc ion binding	8.76251e-10	<0.001	ATP synthesis coupled proton transport
5.44663e-14	<0.001	regulation of nucleobase, nucleoside, nucleotide and nucleic acid metabolic process	9.22263e-10	<0.001	cytoplasmic part
8.03482e-14	<0.001	DNA binding	1.28684e-09	<0.001	ribosome
2.06599e-13	<0.001	regulation of cellular metabolic process	1.54800e-09	<0.001	monovalent inorganic cation transmembrane transporter activity
3.71565e-13	<0.001	regulation of macromolecule biosynthetic process	2.13295e-09	<0.001	translational elongation
6.12201e-13	<0.001	transcription factor activity	5.21561e-09	<0.001	proton transport
6.68339e-13	<0.001	regulation of cellular biosynthetic process	7.45508e-09	<0.001	hydrogen transport
1.06224e-12	<0.001	regulation of biosynthetic process	9.14840e-09	<0.001	ion transmembrane transport
1.45405e-10	<0.001	transcription regulator activity	1.62098e-08	<0.001	structural constituent of ribosome
2.26834e-12	<0.001	regulation of metabolic process	2.19145e-08	<0.001	inorganic cation transmembrane transporter activity
4.89572e-12	<0.001	intracellular	2.42190e-08	<0.001	translation
7.10911e-12	<0.001	regulation of gene expression	3.20020e-08	<0.001	melanosome
1.76818e-09	<0.001	transmembrane receptor protein tyrosine kinase activity	3.20020e-08	<0.001	pigment granule
2.17270e-09	<0.001	transmembrane receptor protein kinase activity	4.99963e-08	<0.001	proton-transporting two-sector ATPase complex, proton-transporting domain
8.56661e-09	<0.001	ephrin receptor activity	7.22247e-08	<0.001	intracellular part
1.19570e-08	<0.001	transmembrane receptor protein tyrosine kinase signaling pathway	5.13700e-08	<0.001	cellular process
1.60773e-07	<0.001	cell differentiation	2.66531e-07	0.001	ATP biosynthetic process

Table 1: The GO enrichment result for cluster 2 identified by DPMMLM.

Table 2: The GO enrichment result for cluster 3 identified by DPMMLM.

P value	Corrected P val	GO term description
2.50635e-12	<0.001	response to wounding
2.20395e-10	<0.001	response to stress
6.78145e-10	<0.001	inflammatory response
7.28686e-10	<0.001	extracellular space
3.81453e-09	<0.001	response to chemical stimulus
6.60775e-09	<0.001	extracellular region part
4.65677e-08	<0.001	response to stimulus
5.76186e-08	<0.001	response to molecule of bacterial origin
1.20234e-07	<0.001	fibrillar collagen
1.37118e-07	<0.001	wound healing
1.88215e-07	<0.001	platelet-derived growth factor binding
2.26731e-07	<0.001	response to organic substance
4.44301e-07	<0.001	response to lipopolysaccharide
6.71932e-07	<0.001	cytokine activity
7.39621e-07	<0.001	response to external stimulus
1.03839e-06	<0.001	developmental process
1.20286e-06	0.001	skin morphogenesis
1.66068e-06	0.001	defense response
1.93814e-06	0.002	receptor binding
2.15753e-06	0.003	chemotaxis

Table 3: The GO enrichment result for cluster 4 identified by DPMMLM.

P value	Corrected P val	GO term description
3.32796e-05	0.048	regulation of receptor biosynthetic process
2.84188e-06	0.007	positive regulation of receptor biosynthetic process

Table 4: The GO enrichment result for cluster 5 identified by DPMMLM.

5 Discussion

If we use a corrected p-value cutoff of 0.001 we see more than 2 fold increase in the number of significant GO categories that are identified by the DPMMLM method compared to the DPMM method (76 vs. 37 with 19 in the overlap). These include some of the most important categories for these experiments as discussed in the main text (for example, apoptosis).

P value	Corrected P val	GO term description
5.66587e-13	<0.001	protein binding
1.76867e-10	<0.001	transcription factor activity
2.23754e-09	<0.001	cellular process
5.84677e-09	<0.001	biological process
9.88355e-09	<0.001	regulation of metabolic process
1.17922e-08	<0.001	regulation of cellular metabolic process
3.15919e-08	<0.001	energy coupled proton transport, down electrochemical gradient
3.15919e-08	<0.001	ATP synthesis coupled proton transport
3.92189e-08	<0.001	regulation of cellular process
7.46999e-08	<0.001	regulation of biological process

P value	Corrected P val	GO term description
7.03858e-16	<0.001	response to stress
6.86006e-13	<0.001	inflammatory response
1.64180e-10	<0.001	immune system process
2.74539e-10	<0.001	immune response
9.16024e-10	<0.001	response to molecule of bacterial origin
4.69286e-09	<0.001	cytokine activity
2.77866e-08	<0.001	receptor binding
3.16405e-08	<0.001	transmembrane receptor protein tyrosine kinase activity
3.46386e-08	<0.001	cytokine receptor binding
4.01148e-08	<0.001	extracellular space
4.87151e-09	<0.001	defense response
5.40508e-09	<0.001	enzyme linked receptor protein signaling pathway
7.83185e-08	<0.001	G-protein-coupled receptor binding
8.13866e-09	<0.001	chemotaxis
8.96811e-09	<0.001	cell communication
1.38699e-07	<0.001	protein binding
1.69750e-07	<0.001	binding
2.24898e-07	<0.001	regulation of transcription, DNA-dependent
3.01156e-07	<0.001	receptor signaling protein activity
3.54444e-07	<0.001	regulation of immune system process

Table 5: The GO enrichment result for cluster 1 identified by DPMM.

Table 6: The GO enrichment result for cluster 2 identified by DPMM.