## More accurate testing via parametric bootstrap

- P-values for MEME/FEL etc are derived from asymptotic approximations (large N)
- Not clear how well these hold for smaller and low-divergence datasets
- Can use a much slower simulation based method to derive more accurate p-values at each site

 Can result both in improved power and lower rates of false positives

## FEL on internal branches of Spike finds most selected sites

Codo	n   Partitio	n   alpha	beta	LRT	Selection detected?
1	1	10000.000	0.075	1.004	Neg. p = 0.0099
5	1	0.000	19.044	2.890	Pos. $p = 0.0297$
12	2 1	0.000	20.323	2.989	Pos. $p = 0.0297$
13	1	0.000	23.877	2.589	Pos. $p = 0.0396$
18	1	0.000	19.110	2.885	Pos. $p = 0.0396$
49	1	0.000	22.768	2.519	Pos. $p = 0.0594$
69	1	0.000	22.573	2.464	Pos. $p = 0.0396$
138	3 1	0.000	26.821	2.736	Pos. $p = 0.0693$
177	1	10000.000	0.000	1.187	Neg. $p = 0.0099$
245	1	0.000	22.499	2.501	Pos. $p = 0.0792$
367	' 1	0.000	44.535	9.044	Pos. $p = 0.0099$
439	1	0.000	34.481	4.986	Pos. $p = 0.0099$
452	2 1	0.000	30.321	5.518	Pos. $p = 0.0099$
477	' 1	0.000	23.678	4.325	Pos. $p = 0.0297$
501	. 1	0.000	38.373	3.317	Pos. $p = 0.0297$
570	) 1	0.000	21.039	3.047	Pos. $p = 0.0594$
614	1	0.000	22.215	3.099	Pos. $p = 0.0594$
681	. 1	0.000	18.344	2.819	Pos. $p = 0.0297$
701	. 1	0.000	20.245	1.998	Pos. $p = 0.0792$
716		0.000	25.450	2.349	Pos. $p = 0.0792$
769	1	0.000	28.107	2.470	Pos. $p = 0.0693$
854	1	0.000	25.908	2.506	Pos. $p = 0.0495$
941	. 1	0.000	25.251	2.344	Pos. $p = 0.0891$
111	.8   1	3.081	27.290	2.023	Pos. $p = 0.0495$
117	76   1	0.000	22.141	3.039	Pos. $p = 0.0297$
123	37   1	10000.000	0.151	0.876	Neg. $p = 0.0099$
124	8   1	0.000	20.075	2.336	Pos. $p = 0.0891$

## CAUTION: A VERY TIME CONSUMING ANALYSIS (SEVERAL HOURS)

hyphy fel --alignment data/spike.fas --tree data/spike.tree --branches Internal --output Spike-pbs.FEL.json --resample 100