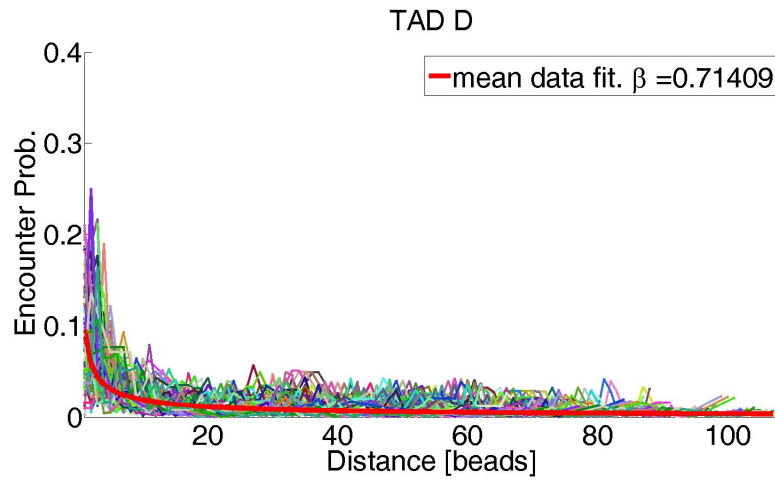


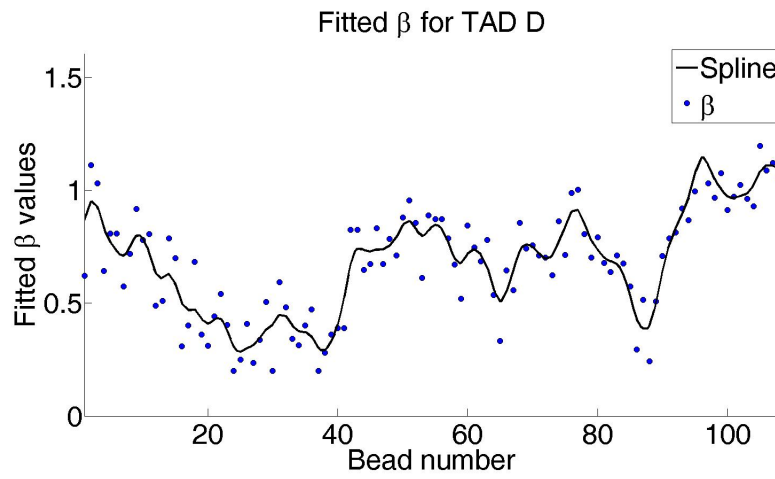
## 0.1 Peak Calling

To accurately identify pair of beads that interact more frequently than expected, we ran a peak calling procedure on the 5C probability data of each TAD separately. For each bead  $i = 1..N$  of the chain, we fit an encounter probability curve of the form  $\alpha_i d^{-\beta_i}$ . We then use the values of the fitted curves at each distance  $d = 1..(N - 1)$  to estimate the expected encounter probability,  $\mu(d) = E[\alpha_i d^{-\beta_i}]$ , and the expected standard-deviation,  $\sigma(d) = \sqrt{E[(\mu(d) - \alpha_i d^{-\beta_i})^2]}$ , for each  $d$ , and  $i = 1..N$ .

For each observation  $i$  at distance  $d$  we calculate a z-score by  $z_i(d) = \frac{|\alpha_i d^{-\beta_i} - \mu(d)|}{\sigma(d)}$  and fit a Weibull distribution to it. For each z-score a p-value is calculated based on the fitted Weibull CDF and then transformed into a q-value to obtain false discovery rate. We then threshold the data using false discovery rate of 0.01.

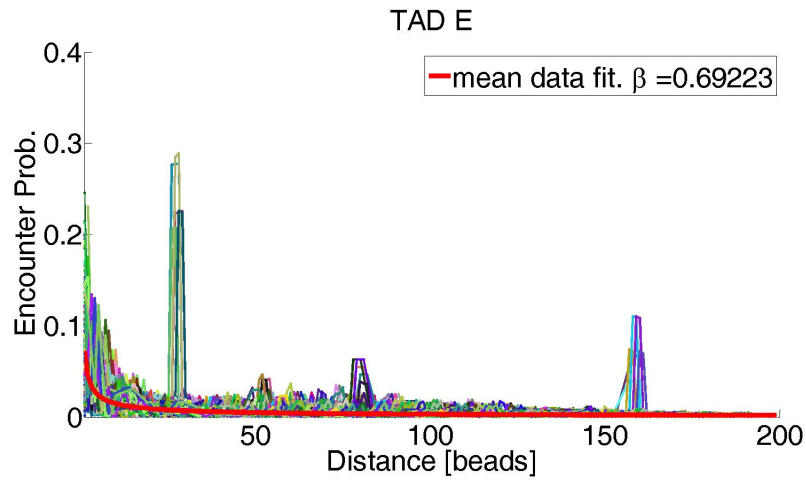


(a)

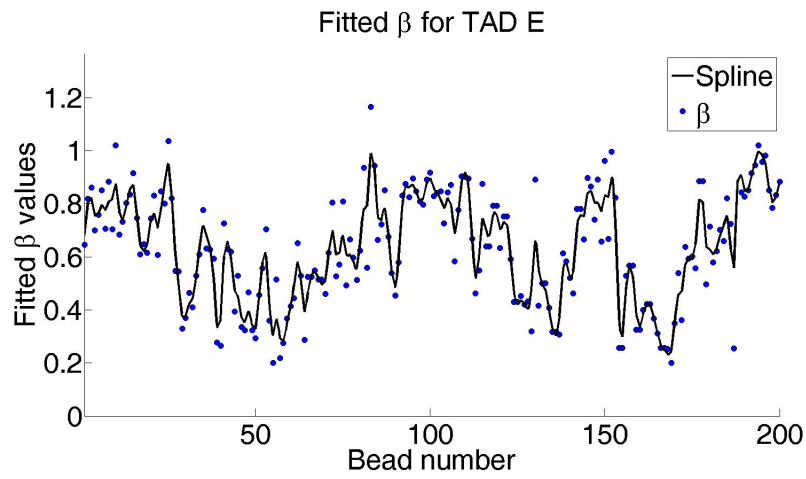


(b)

Figure 1: The encounter probability and the fitted  $\beta$  values for TAD D.



(a)



(b)

Figure 2: The encounter probability and the fitted  $\beta$  values for TAD E.