

MATH-578B: Assignment # 3

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Problem 1

The coverage c depends on the position x as: $c = \frac{NL_x}{G}$ where L_x is the expected length of clones covering x .

Probability any position x to be covered by atleast one clone = $(1 - \text{Probability that it is sequenced by atleast one clone})$.

Probability that position x is not sequenced = Probability of zero clones starting in $(x - L, x]$ = No arrivals in the interval $(x - L, x] = e^{-c(x)}$

Probability that it is sequenced = $1 - e^{-c(x)}$ where $c(x)$ represents that c is a function of x .

$C \sim \Gamma(\alpha, \beta)$

$$f(c) = \frac{c^{\alpha-1} e^{-c/\beta}}{\beta^\alpha \Gamma(\alpha)}$$

Thus,

$$P(N_h = k) = \int_0^\infty e^{-ch} \frac{(ch)^k}{k!} \times \frac{c^{\alpha-1} e^{-c/\beta}}{\beta^\alpha \Gamma(\alpha)} dc$$

Problem 2

Given: $\lim_{n \rightarrow \infty} (1 - F(b \log(n) + x/a)) = G(x)$

$$\lim_{n \rightarrow \infty} (1 - F(b \log(n) + x/a)) = G(x)$$

$$\lim_{n \rightarrow \infty} F(b \log(n) + x/a) = 1 - G(x)/n$$

$$\begin{aligned} P(a(\max_i X_i - b \log(n)) \leq x) &= P(\max_i X_i \leq x/a + b \log(n)) \\ &= P(X_1 \leq x/a + b \log(n)) P(X_2 \leq x/a + b \log(n)) \dots P(X_n \leq x/a + b \log(n)) \\ &= (F(x/a + b \log(n)))^n \\ &= \lim_{n \rightarrow \infty} n \log(1 - G(x)/n) \\ &= \lim_{n \rightarrow \infty} e^{n \log(1 - G(x)/n)} \\ &= e^{-G(x)} \end{aligned}$$

Choosing a, b for $G(x) = e^{-x}$ given $X_i \sim \text{exponential}(\lambda)$

$$f(x|\lambda) = \lambda e^{-\lambda x} \implies F(x) = 1 - e^{-\lambda x}$$

Now,

$$\begin{aligned} \lim_{n \rightarrow \infty} 1 - G(x)/n &= F(b \log(n) + x/a) \\ &= 1 - e^{-\lambda(b \log(n) + x/a)} \\ e^{-x}/n &= e^{-\lambda(b \log(n) + x/a)} \\ -x &= \log(n) + -\lambda(b \log(n) + x/a) \\ x(-1 + \lambda/a) &= \log(n) - b\lambda \log(n) \end{aligned}$$

Thus,

$$a = \lambda; b = \frac{1}{\lambda}$$

Problem 3

Target Distribution in aligned region: $P(R, R) = 0.2$; $P(Y, Y) = 0.7$; $P(R, Y) = 0.1$

$$\xi_r = 0.2$$

$$\xi_y = 0.8$$

By Theorem 11.7 we have that the proportion of letter a aligning with letter b in the best matching interval converges to:

$$p(a, b) = \xi_a \xi_b p^{-s(a, b)}$$

Equivalently:

$$s(a, b) = \log_{1/p} \left(\frac{p(a, b)}{\xi_a \xi_b} \right)$$

$$p = \xi_r \xi_r + \xi_y \xi_y = 0.68$$

Thus

$$P(RR) = \xi_r \xi_r p^{-s(r, r)}$$

$$\begin{aligned} s(r, r) &= \log_{1/0.68} \left(\frac{0.2}{0.04} \right) \\ &= 4.17 \end{aligned}$$

$$s(r, y) = \log_{1/p} \left(\frac{p(r, y)}{\xi_r \xi_y} \right)$$

$$\begin{aligned} s(r, y) &= \log_{1/0.68} \left(\frac{0.1}{0.16} \right) \\ &= -1.21 \end{aligned}$$

$$s(y, y) = \log_{1/p} \left(\frac{p(y, y)}{\xi_y \xi_y} \right)$$

$$\begin{aligned} s(y, y) &= \log_{1/0.68} \left(\frac{0.7}{0.64} \right) \\ &= 0.23 \end{aligned}$$

$s(r, r) = 4.17$

$s(y, y) = 0.23$

$s(y, r) = -1.21$

To find the value of λ such that:

$$\lim_{n \rightarrow \infty, m \rightarrow \infty} P\{\lambda R_{mn} - \log(K_{mn}) < x\} = \exp(-\exp(-x))$$

$$\lambda = \log(1/p) = 0.38$$

And given that the score for 1000bp alignment is 100, the p value is given by;

$$p - \text{value} = 1 - e^{-e^{-s}}$$

where $s = \lambda R_{mn} - \log(K_{mn})$

If the p-value is less than some pre-defined threshold, the hypothesis that alignment is as good as by random chance can be rejected.

Problem 4

Minimal neighborhood set $J_{i,j}$ such that $\{i', j' \in J_{i',j'}^c\}$ are independent of $Y_{i,j}$ is given by: $\{(i', j') : |i - i'| \leq t \text{ or } |j - j'| \leq t\}$

Now,

$$\begin{aligned} b_1 &= \sum_{i \in I} \sum_{j \in J_i} E(X_i) E(X_j) \\ &= p^t \sum_{j \in J_i} E(X_j) + \sum_{i=2}^{n-t+1} (1-p)p^t \sum_{j \in J_i} E(X_j) \\ &= (n-t+1)p^t(2t+1)p^t \times 2 + (n-t+1)^2(1-p)^2 p^{2t}(4t+2) \\ &= p^{2t}(n-t+1)(4t+2)(1 + (n-t+1)(1-p)^2) \end{aligned}$$

$$\begin{aligned} E[NC_n] &= (n-t+1)(1-p)p^t \\ &= \lambda \end{aligned}$$

Thus, $n(1-p)p^t \sim \lambda$

$$\log_{1/p} \lambda = \log_{1/p} n +$$

Choose $t_n = 2 \log_{1/p} n + x$

Consider $(n - t_n + 1)p^{2t} = (n - \log_{1/p} n^2 + 1)p^{2t}$

Exercise 11.9

Part (a): $\xi_a = 1/|A|$

$$p = \sum_{a \in A} \xi_a \xi_a = |A|/|A|^2 = 1/|A|$$

$$\begin{aligned} \mu_{a,a} &= \xi_a \xi_a p^{-s(a,a)} \\ &= 1/|A|^2 \times |A| \\ &= 1/|A| \end{aligned}$$

Part (b): To Derive $s(a, b)$ such that $\mu_{a,a} = 1/|A|$ We have (from problem 3):

$$s(a, b) = \log_{1/p} \left(\frac{p(a, b)}{\xi_a \xi_b} \right)$$

And hence:

$$s(a, a) = \log_{1/p} \left(\frac{1/|A|}{1/|A|^2} \right) = 1$$

Thus,

$$s(a, a) = 1; s(a, b) = -\infty$$

Exercise 11.11

If there are N sequences we have n^N choices for $(i_1, i_2, i_3 \dots i_n)$ where i_j represents the starting position for the j^{th} sequence. and hence using naive approach (Largest run being unique and satisfying $np^{R_n} = 1$ and then by allowing shifts there are n^N choices for starting position in each sequence):

H_n grows like

$$\log_{1/p} n^N = N \log_{1/p} n$$

and hence

$$t(n, N) = N \log_{1/p} n$$

As

$$N \rightarrow \infty \implies t(n, N) \rightarrow \infty$$