

Though we have a DP algorithm to compute the hit probability of a seed, it is still useful to have some sense of the “dependency” between the 1’s and its relation with the general hit probability. In the following problem, consider two seeds:

Seed_X = 1101011

Seed_Y = 11111

where

1 for match and 0 for don’t care in the seeds

L = the length of the query and subject sequence

p = similarity of the two sequence = Prob(the two sequence has same symbol)

M = the length of the seed

W = the weight of the seed

assuming that there are only match/mismatch in any pair of symbols and overlapped hits counts as different hits

1. In the two seeds, which seed is likely to be used in PatternHunter?

A: Seed_X since PatternHunter uses spaced seeds

2. If we already have a hit in a pair of sequence, what’s the expected total number of hits we can find in this pair?

A: If we have already a match in the positions of 1’s of a hit, we need less new matches than its actual weight in the region overlapping the first hit. We can regard Seed_Y as “1111100” which has the same window length as Seed_X does, and in the overlapped regions the probability to find a second hit is

For Seed_X: $(p^3 + p^3 + p^3 + p^4 + p^3 + p^4) * 2$

For Seed_Y: $(p + p^2 + p^3 + p^4 + p^5 + p^5) * 2$

the probability is multiplied by 2 since the window can be shifted backwards and forwards.

Outside the region, the probability to find a second hit is independent of the first hit, hence it is p^5 for all possible positions for both seeds.

3. If we can approximate the probability to hit by the formula

$E(\#hits) = Pr(seed\ hits) * E(\#hits | seed\ hits)$

what’s the order of the seeds in terms of E(#hits)? in terms of Pr(seed hits)? which criterion is more important, why?

A: $E(\#hits)$ are $(L - M + 1) p^W$ for both seeds, hence $E_X(\#hits) = E_Y(\#hits)$

$Pr(seed\ hits) = E(\#hits) / E(\#hits | seed\ hits)$, hence $Pr_X(seed\ hits) > Pr_Y(seed\ hits)$

$Pr(seed\ hits)$ is more important, since ideally we need just one hit for further extension in each significant alignment of two homologous sequences; in other

words, finding more hits in the same alignment does not contribute to the sensitivity of the alignment algorithm.