

PLNT4610 BIOINFORMATICS

FINAL EXAMINATION

9:00 to 11:00 Friday December 6, 2013

Answer any combination of questions totalling to exactly 100 points. The questions on the exam sheet total to 120 points. If you answer questions totalling more than 100 points, answers will be discarded at random until the total points equal 100. This exam is worth 20% of the course grade.

Hand in the question sheets along with your exam booklet. All questions must be answered in the exam book. The question sheets will be shredded after the exam.

1. (10 points) When a multiple sequence alignment contains large gaps, what problem does this create for constructing phylogenetic trees? What is one solution to constructing phylogenies, when the alignment contains large gaps?

2. (10 points) Given the following list of terms, draw a DAG (directed acyclic graph) that describes an ontology for the following terms:

Terms: Maximum likelihood; phylogeny methods; probabilistic methods; Fitch least squares; Bayesian phylogeny; Character methods; Neighbor joining; Distance methods; parsimony

3. (5 points) Many phylogenetic analysis programs have an option to jumble the order of sequences. What is the reason for this function, and what does it accomplish?

4. (10 points) The table below shows the relative frequencies with which different types of molecular markers change. Explain the reasons behind these relative frequencies.

RFLP	$1 \rightarrow 0 > 0 \rightarrow 1$
RAPD, AFLP, SRAP	$1 \rightarrow 0 \gg 0 \rightarrow 1$
microsatellites	$1 \rightarrow 0 = 0 \rightarrow 1$

5. (10 points) List five sources of experimental variance in microarray experiments. Use point form for your answer.

6. (10 points) The equation at right is used for screening molecular markers for linkage to a Mendelian trait. State the meaning of the terms N, P and f.

$$N = \frac{\ln(1-P)}{\ln(1-f)}$$

7. (5 points) Why can't we say that genetic distances on a chromosome are directly proportional to physical distances?

8. (15 points) In a cross between two *Arabidopsis* lines, A and B, a map of one chromosome was constructed using a set of co-dominant markers. An excerpt of the mapping data for this cross is shown in panel I. At each locus, the marker is scored as being homozygous for the allele from parent A, homozygous for the allele from parent B, or heterozygous. The order of loci shown in the table corresponds to the order of those loci on the chromosome.

a) What is the predicted ratio for seeing A, H or B, at any given locus?

b) In cross II, parent A was crossed with another *Arabidopsis* line, C. Thus, the expected phenotypes would be either A, H or C. In this cross, the mapping data look similar to that found in cross I. However, all loci distal to g3883 exhibit only the A phenotype, in all progeny. What is a simple explanation for this result?

c) Based on your answer to b, how could you test your hypothesis?

I. A x B		II. A x C	
segregating progeny ----->		segregating progeny ----->	
marker/ map posn.		marker/ map posn.	
g6844	HHA AAAAABVHHBAAAHVHHHHB VHHHHB VBAHVBH AHHHVA AHNA	g6844	HHA AAAAACHHCAAHCHHHHAC HHHAC SAHHC AHHHCA AHHA
g3843	HHA AAAAABVHHBAAAHVHHHHB VHHHHB VBAHVBH AHHHVA AHNA	g3843	HHA AAAAACHHCAAHCHHHHAC HHHHAC HHAH CAHHC AHHHCA CA
g2616	HHA AHVHHVHHBAAAHVHHHHB VHHHHH HVBVHHA AHHHHHHH H	g2616	HHA AHHCCHHCCAAHCHHHHAC HHHHHHC SC HCHHA HHHHHH H
m210	HHA AHVHHHHHAAAAHVHHHHAN HAAHNAH HHA ABHHAHVBAA	m210	HHA AHCHHHHHAAAAHCHHHHAAHNA HAAHNAH HHA ACHHAAHCA
g6837	HHA ABVHHAVHHBAAAHVHHHAAHNAH HHA AVHHAVHBA	g6837	HHA ACHHAAHCHHCAAHCHHHHAAHNAH HHA ACHHAAHCHCA
g10086	AHNA AHVHHHAVHHVBAHHHHAAHNAH HHA AHVHHHVB	g10086	AHNA AHVHHHAAHCHHCAAHCHHHHAAHNAH HHA AHVHHHCHC
g4564a	HA AHVHHHHHAAAAHVHHHAAHNAH HHA AHVHHHVA	g4564a	HA AHVHHHHHAAAAHCHHHHAAHNAH HHA AHVHHHAAHCHHCA
g3845	HA HHHVHHHAAAAHVHHAAHVAAH HHA AHVHHHVA	g3845	HA HHHCHHHHAAAAHCHHAAHCAH HHA AHVHHHAAHCHHAAH
g4539	AHNA AHVHHHAAHVAHHAHNAH HHA HAAHNAH HVAH VHHHAAH	g4539	AHNA AHVHHHAAHCHHAAHCAH HHA HAAHNAH HAAHCHHAAH
m557	HA HHHVHHHAAAAHVHHAAHVAH HHA AHVHHHVAH VHHHAAH	m557	HA HHHCHHHHAAAAHCHHAAHCAH HHA AHVHHHAAHCHHAAH
g3883	HA HHHVHHHAAAAHVHHHAAHVAH HHA AHVHHHVAH VHHHAAH	g3883	HA HHHCHHHHAAAAHCHHAAHCAH HHA AHVHHHAAHCHHAAH
g19833	HA HAVHHHAAAAHVHHAAHVAH HHA AHVHHHVAH VBAH VBAH	g19833	AA
g19838	HA HAVHHHAAHVAHAAHVAH HHA AHVHHAAHVAH HHA HVAH HHAH	g19838	AA
m272	HA HAVHHHAAHVAHAAHVAH HHA AHVHHAAHVAH HHA HVAH HHAH	m272	AA
g4513	HA HAVHHHAAAAHVHHHHBAAAHVHHHAAHVAH VBAH HHHAV	g4513	AA

9. (10 points) What is the basis for polymorphism with Microsatellite markers? Why are microsatellite markers usually more informative than markers such as RFLPs, RAPDs or SCARS, that only have 2 possible states?

10. (5 points) What is meant by the term "homoplasy", and why is it important in phylogenetic analysis?

11. (10 points) A series of random DNA sequences was constructed, each with a different percentage of AT bases. and sequences were compared using several phylogeny methods. In each case, 100 bootstrap replicates were done. The results are presented in the table below.

ran20 - 20% AT ran35 - 35% AT ran50 - 50% AT ran65 - 65% AT ran80 - 80% AT	
Neighbor Joining <pre> +-----ranAT65 +--42.0- +-----ranAT80 +----- +-----ranAT20 +--32.0- +-----ranAT35 +-----ranAT50 </pre>	Fitch/Margoliash <pre> +-----ranAT35 +--24.0- +-----ranAT20 +----- +-----ranAT80 +-----ranAT50 +-----ranAT65 </pre>
Parsimony <pre> +-----ranAT35 +----- +-----ranAT50 +--100.0- +-----ranAT80 +--100.0- +-----ranAT65 +-----ranAT20 </pre>	Maximum Likelihood <pre> +-----ranAT20 +--62.0- +-----ranAT35 +----- +-----ranAT50 +-----ranAT80 +-----ranAT65 </pre>

- First, consider the parsimony tree. Explain why the sequences group as they do in this tree.
- Can you think of a reason why the distance methods showed lower bootstrap results than did parsimony and maximum likelihood?

12. (10 points) Describe the process of bootstrap resampling, as applied to phylogenetic analysis of DNA or protein sequences.

13. (10 points) A BLASTP hit is shown below in two formats. The first is the familiar report format, showing the alignment between the query sequence and a matching sequence in the Patented division of GenBank. The second is the corresponding XML output produced by BLASTP.

Based on the XML, draw a database schema diagram for a BLAST hit. You can assume that Hit and Hsp are two distinct classes. The Hit class would have a field called Hit_hsps, which points to a list of objects of the class Hsp.

REMEMBER: You are being asked to create classes, not objects.

```
>emb|CAA01678.1| acidic chitinase SE [Beta vulgaris subsp. vulgaris]
Length=293
```

```
Score = 339 bits (869), Expect = 3e-115, Method: Compositional matrix adjust.
Identities = 171/253 (68%), Positives = 199/253 (79%), Gaps = 4/253 (2%)
```

```
Query 1 IAVYWQNGGEGSLADTCNTGNYEFVNI AFLSTFGSGQTPQLNL LAGHCDPSSNGCTGFSS 60
      I +YWGQNG EGSLADTCN+GNY V +AF++TFG+GQTP LNL LAGHCDP++N C SS
Sbjct 28 IVIYWQNGGEGSLADTCNSGNYGTVILAFVATFGNGQTPALNL LAGHCDPATN-CNSLSS 86

Query 61 EIQTCQNRGIKVL LSLGGSAGTYSLNSADDATQLANYLWDN FLGGQSGSRPLGDAVL DGV 120
      +I+TCQ GIKVLLS+GG AG YSL+S DDA A+YLW+ +LGGQS +RPLGDAVL DG+
Sbjct 87 DIKTCQQAGIKVLLS IGGGAGGYSLSSTDDANTFADYLWNTY LGGQSSTRPLGDAVL DGI 146

Query 121 DFDIESGGSNHYYDDLARALNSLSS-QKKVYLSAAPQC IIPDQHLDAAIQTGLFDYVWVQF 179
      DFDIESG +DDLARAL ++ QK VYLSAAPQC +PD L AI TGLFDYVWVQF
Sbjct 147 DFDIESGDRGFWDDLARALAGHNNGQKTVYLSAAPQCPLPDASLSTAIATGLFDYVWVQF 206

Query 180 YNNPSCQYNSGGTTNLINSWNQWITVPASLVFMGLPASDAAAPSGGFVSTDVLT SQVLPV 239
      YNNP CQY NL++SWNQW TV A+ +F+GLPAS AA S GF+ D LTSQVLP
Sbjct 207 YNNPPCQYDTSA-DNLLSSWNQWTTVQANQIFLGLPASTDAAGS-GFIPADALTSQVLP T 264

Query 240 IKQSSKYGGVMLW 252
      IK S+KYGGVMLW
Sbjct 265 IKGSAKYGGVMLW 277
```

```
<Hit>
  <Hit_num>5</Hit_num>
  <Hit_id>gi|904330|emb|CAA01678.1|</Hit_id>
  <Hit_def>acidic chitinase SE [Beta vulgaris subsp. vulgaris]</Hit_def>
  <Hit_accession>CAA01678</Hit_accession>
  <Hit_len>293</Hit_len>
  <Hit_hsps>
    <Hsp>
      <Hsp_num>1</Hsp_num>
      <Hsp_bit-score>339.347</Hsp_bit-score>
      <Hsp_score>869</Hsp_score>
      <Hsp_evalue>3.28347e-115</Hsp_evalue>
```

```

<Hsp_query-from>1</Hsp_query-from>
<Hsp_query-to>252</Hsp_query-to>
<Hsp_hit-from>28</Hsp_hit-from>
<Hsp_hit-to>277</Hsp_hit-to>
<Hsp_query-frame>0</Hsp_query-frame>
<Hsp_hit-frame>0</Hsp_hit-frame>
<Hsp_identity>171</Hsp_identity>
<Hsp_positive>199</Hsp_positive>
<Hsp_gaps>4</Hsp_gaps>
<Hsp_align-len>253</Hsp_align-len>
<Hsp_qseq>IAVYWGQNGGEGSLADTCNTGNYEFVNIAFLSTFGSGQTPQLNLAGHCDPSSNGCTGFSSEIQCQN
RGIKVLLSLGGSAGTYSLSNADDATQLANYLWDNFLGGQSGSRPLGDAVLGDVDFDIESGGSNHYDDLARALNSLSS-
QKKVYLSAAPQCIIPDQHLDAAIQTGLFDYVWVQFYNNPSCQYSNGGTTNLINSWNQWITVPASLVFMGLPASDAAAPSGGFVSTDVLT SQ
VLPVIKQSSKYGGVMLW</Hsp_qseq>
<Hsp_hseq>IVIIYWGQNGDEGSLADTCNSGNYGTVILAFVATFGNGQTPALNLAGHCDPATN-
CNSLSSDIKTCQQAGIKVLLSIGGGAGGYSLSSTDDANTFADYLWNTYLGGQSSTRPLGDAVLGDGIDFDIESGDGRFWDDLARALAGHNNG
QKTVYLSAAPQCPLPDASLSTAIATGLFDYVWVQFYNNPPCQYDTSA-DNLLSSWNQWTTVQANQIFLGLPASTDAAGS-
GFIPADALTSQVLPTIKGSAKYGGVMLW</Hsp_hseq>
<Hsp_midline>I +YWGQNG EGSLADTCN+GNY V +AF++TFG+GQTP LNLAGHCDP++N C
SS+I+TCQ GIKVLLS+GG AG YSL+S DDA A+YLW+ +LGGQS +RPLGDAVLGDG+DFDIESG +DDLARAL ++ QK
VYLSAAPQC +PD L AI TGLFDYVWVQFYNNP CQY NL++SWNQW TV A+ +F+GLPAS AA S GF+ D LTSQVLP
IK S+KYGGVMLW</Hsp_midline>
</Hsp>
</Hit_hsps>
</Hit>

```