Intro. to Graphical Models and RevBayes



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What is a **model**?



Equations and plot: https://en.wikipedia.org/wiki/Lotka-Volterra_equations

What is a **model**?

Mathematical models try to explain important features of (biological) systems using (relatively simple) mathematical principles.



Equations and plot: https://en.wikipedia.org/wiki/Lotka–Volterra_equations

Lynx photo: Wikipedia (NPS Photo/Jacob W. Frank) Snowshoe Hare photo: Wikipedia (D. Gordon E. Robertson)

Deterministic and Stochastic Models

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NO randomness

(no stochasticity) (no probability statements)

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Randomness!

(stochasticity) (probability statements) (variation in possible outcomes)

Stochastic Models



Randomness!

(stochasticity) (probability statements) (variation in possible outcomes)

A simple stochastic model



Bernoulli

Describes the outcome of a single coin flip (or other binary 0/1 trial)

A simple stochastic model



Bernoulli One type of *discrete* distribution

A simple stochastic model



P(Heads) = pP(Tails) = 1-p

Let's build this model!



RevBayes



moleuser@amphioxis-49-of-66:~\$ rb

RevBayes version (1.1.1) Build from tags/1.1.1 (rapture-588-gae00cc) on Sat Mar 19 00:48:15 UTC 2022 Visit the website www.RevBayes.com for more information about RevBayes. RevBayes is free software released under the GPL license, version 3. Type 'license()' for details.

To quit RevBayes type 'quit()' or 'q()'.

https://revbayes.github.io

Let's build this stochastic model!



Ζ

Let's build this stochastic model!





```
dnBernoulli(0.5)
Z
       dnBernoulli(0.5)
Z
       dnBernoulli(0.5)
       dnBernoulli(0.5)
       dnBernoulli(0.5)
       dnBernoulli(0.5)
Z
       dnBernoulli(0.5)
Z
 0
```

When you create a random variable in RevBayes, it automatically initializes that variable by drawing a value from the corresponding probability distribution.

The Role of Models

(1) Can we explain the **important features** of the real world using relatively **simple mathematical principles** (prediction)?

(2) Within this mathematical framework, can we **learn about the processes** that generated our data (inference)?

The most useful models can answer 'yes' to both.

$$z \sim dnBernoulli(0.5)$$

 z
(1)
(2)
 $p=0.5$
No data

Different parameter values imply different probabilities for outcomes (different PMFs)





Different parameter values imply different probabilities for outcomes (different PMFs)



Computing Likelihoods







> p <- 0.75
> z ~ dnBernoulli(p)
> z.clamp(1)
> z.probability()
0.75

$$p = 0.25$$
 $p = 0.5$ $p = 0.75$





Graphical Models

Graphical models provide a means of depicting the dependencies among parameters in probabilistic models.



The "graphical" part of the name graphical models has to do with their basis on graphs, and not anything to do with drawing.

But it is convenient that graphical models can also be drawn, to clearly depict the structure of the model.

Building a Graphical Model

p is fixed at 0.25



<- 0.25 D

z depends on *p z* is a Bernoulli r.v.

We "observed" a value of 1 for *z*.



> z ~ dnBernoulli(p)





What are the possible building blocks for graphical models?



RevBayes Syntax

sqrt(16) abs(-11)	functions and arguments	choose(6,3) ls() max([2.1,5.5,7.8])
test <- v(5,9,12) test test[2]	vectors	test[1] <- 5 test[2] <- 9 test[3] <- 12 test
for (i in 1:5){		for (i in 1:3){

https://revbayes.github.io/tutorials/intro/rev.html https://revbayes.github.io/documentation/

for loops

print(test[i])

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print(i)



Type Constant Node

Description

Constant nodes are like standard variables in a programming language. They are assigned fixed values.

Rev Example

x <- 2.3



Type Deterministic Node

Description

Deterministic nodes depend on the values of other nodes, but in a deterministic (fixed) way. They have no probability distribution intrinsically associated with them.

Rev Example



Type Stochastic Node

Description

Stochastic nodes represent random variables and take on values according to some probability distribution. These distributions may have parameters defined in other nodes in the model graph. The type of distribution associated with a node is often not written explicitly when a model is drawn, but it must be specified.

Rev Example

 $z \sim dnBernoulli(0.5)$





Type Clamped Stochastic Node

Description

Data can be viewed as the observed outcome of a stochastic node. Clamping involves assigning a set of observations to an associated stochastic node. Clamping data allows the values of other parameters in the model to be inferred.

Rev Example

z ~ dnBernoulli(0.5) z.clamp(1)

Symbol



Type Conditional Dependency

Description

Arrows indicate conditional dependencies and are directional. The node (variable) that the arrow points to depends on the value of the node at the other end of the arrow. For instance, if *p* points to *Z*, then we say that *Z* is dependent on *p*.

Rev Example

p <- 0.5z ~ dnBernoulli(p) s := z*3

$$\boxed{0.5} \longrightarrow \overrightarrow{Z} \longrightarrow \overrightarrow{S}$$



Description

Plates simply represent repetition and make it easier to depict graphical models with a repetitive structure. For instance, the example above shows *N* instances of the clamped variable, *z*, with each instance indexed by *i*. Plates can be used with any type of node, but are commonly used with clamped nodes representing data.

Rev Example



What's the structure of this model? What would happen if *x* was set to 16? **Try changing** *x* **without changing** *y***, but then look at value of** *y***.**





What's the structure of this model? What can we infer about this model?





What's the structure of this model? What can we infer about this model? **Nothing to infer! Only nodes are constant or clamped.**





What's the probability of our "observed" value?
Probability Density Function (PDF)



Single Bernoulli Trial



What's the structure of this model? What can we infer about this model? How much information will we have?

Single Bernoulli Trial



p ~ dnUnif(0,1) x ~ dnBernoulli(p) x.clamp(0)

Series of Bernoullis with Linked p



What can we infer about this model? How much information will we have (relative to a single Bernoulli)?

Series of Bernoullis with Linked p



Single Binomial



How does this model relate to the Bernoulli models?

Single Binomial





First, create a model object. You can pass any of the nodes to the constructor as a "handle".

myModel = model(n)

NOTE: We use the = assignment operator for "workspace" variables.



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Next, we need to define a proposal distribution (move) for any parameters we are trying to infer.

moves = VectorMoves()
moves.append(mvSlide(p,delta=0.1,weight=1))

Many different move types are available in RevBayes.

As our MCMC runs, we need to keep track of our progress and sampled parameter values. To do that we use monitors.

monitors = VectorMonitors() monitors.append(mnScreen(printgen=1000,p)) monitors.append(mnModel(filename="myMCMC.log", printgen=10))

Running MCMC simulation This simulation runs 1 independent replicate. The simulator uses 1 different moves in a random move schedule with 1 moves per iteration

Iter	I	Posterior	Ι	Likelihood	Ι	Prior	I	р	Ι	elapsed	Ι	ETA	Ι
0	 I	-10.2115	 I	-10.2115		0		0.08309243	 	00:00:00		::	
1000	i	-4.8278	i	-4.8278	i	0	i	0.5133643	i	00:00:00	i	::	i.
2000	i	-4.97547	i	-4.97547	i	0	i	0.453571	i	00:00:00	i	00:00:00	i
3000	i	-4.78164	i	-4.78164	i	0	i	0.5619349	i	00:00:00	i	00:00:00	i
4000	i	-5.18177	i	-5.18177	i	0	i	0.7290954	i	00:00:00	i	00:00:00	i
5000	i	-5.98066	i	-5.98066	i	0	i	0.2898232	i	00:00:00	i	00:00:00	i
6000	i	-4.78464	i	-4.78464	i	0	i	0.5540578	i	00:00:00	i	00:00:00	i
7000	i	-4.78044	i	-4.78044	i	0	i	0.5690397	i	00:00:00	i	00:00:00	i
8000	i	-4.79273	i	-4.79273	i	0	i	0.541859	i	00:00:00	i	00:00:00	i
9000	i	-4.96352	i	-4.96352	i	0	i	0.4572184	i	00:00:00	i	00:00:00	i.
10000	i	-4.80438	Ì	-4.80438	Ì	0	İ	0.6120314	İ	00:00:00	Í	00:00:00	İ

Next we create an MCMC object.

myMCMC = mcmc(myModel,moves,monitors)

Now, we start the MCMC!

myMCMC.run(20000)

Wait - what does this 20,000 mean?

Tracer



Marginal Distribution

Trace Plot

Assessing Convergence



Assessing Convergence



How many samples did we draw during our MCMC analyses?

95% Highest Posterior Density (HPD) Intervals



Phylogenetic Graphical Models

Model Space (conceptual)

All Possible Ways Genes Could Evolve

Model Space (conceptual)

All Possible Ways Genes Could Evolve



Our Hope

All Possible Ways Genes Could Evolve



Our Fears

All Possible Ways Genes Could Evolve



The Probable Truth

All Possible Ways Genes Could Evolve



The Probable Truth

All Possible Ways Genes Could Evolve



The Types of Evolution That Are Built Into Many Programs

Types of Evolution that You'd Like to Try!







data = readDiscreteCharacterData("myData.nex") taxa <- data.taxa() n_taxa <- data.ntaxa() n_branches <- 2 * n_taxa - 3

topology ~ dnUniformTopology(taxa)

alpha <- 2 beta <- 4

TL ~ dnGamma(alpha,beta)

alpha_bl <- 1.0

rel_branch_lengths ~ dnDirichlet(rep(alpha_bl,n_branches))

br_lens := rel_branch_lengths * TL

Q <- fnJC(4)

psi := treeAssembly(topology, br_lens)

seq ~ dnPhyloCTMC(tree=psi,Q=Q,type="DNA")



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Jukes-Cantor

(Compound Dirichlet tree- and branch-length prior)



Tree Length $Gamma(\alpha, \beta)$

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Jukes-Cantor



Inferring these parameters under Jukes-Cantor.







How does HKY85 differ from JC?





How does HKY85 differ from JC?

Transition-Transversion Rate Ratio

mu <- 0 sigma <- 1 kappa ~ dnLognormal(mu,sigma)

alpha_pi <- 1.0 pi ~ dnDirichlet(rep(alpha_pi,4))

Q := fnHKY(kappa,pi)





How does HKY85 differ from JC?

mu <- 0 sigma <- 1 kappa ~ dnLognormal(mu,sigma)

alpha_pi <- 1.0 pi ~ dnDirichlet(rep(alpha_pi,4))

Variable Base Frequencies

Q := fnHKY(kappa,pi)



General Time Reversible (GTR)

Variable Exchangeabilities

alpha_er <- 1.0 er ~ dnDirichlet(rep(alpha_er,6))

alpha_pi <- 1.0 pi ~ dnDirichlet(rep(alpha_pi,4))

Q := fnGTR(er,pi)



Things to Remember

As we add more stochastic nodes, **remember to assign moves** to all of them!

Vectors modeled with a Dirichlet should have a joint proposal mechanism, since they need to sum to 1. Here are a couple:

mvBetaSimplex mvDirichletSimplex

You'll often want to run analyses on the same dataset with multiple models. Start by saving your simplest model into a text file (.rev). Then, copy and paste the code into a file for the next model and adjust as necessary. Keep going as you work your way to more complex models.

Topology Moves



Nearest-Neighbor Interchange (NNI)

The most minor topological move.

https://en.wikipedia.org/wiki/Tree_rearrangement

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Topology Moves



Subtree Prune and Regraft (SPR)

Intermediate topological move.

Often a good balance between exploring new tree space without disrupting things too much.

https://en.wikipedia.org/wiki/Tree_rearrangement

Topology Moves



Tree Bisection and Reconnection (TBR)

Disruptive topological move.

https://en.wikipedia.org/wiki/Tree_rearrangement



Modeling Rate Variation Across Sites Gamma Distribution

mu_a <- ln(5.0) sigma_a <- 0.587405

alpha ~ dnLognormal(mu_a, sigma_a)

sr := fnDiscretizeGamma(alpha, alpha, 4, false)

seq ~ dnPhyloCTMC(tree=psi,Q=Q, siteRates=sr,type="DNA")



LogNormal

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Can add + Γ to any "standard" model of

sequence evolution.

seq

Discretized Gamma

Modeling Rate Variation Across Sites **Proportion of Invariable Sites**





Modeling Rate Variation Across Sites Proportion of Invariable Sites

alpha_p <- 1.0 beta_p <- 1.0

```
p_inv ~ dnBeta( alpha_p, beta_p )
```

```
seq ~ dnPhyloCTMC(tree=psi, Q=Q,
plnv=p_inv, type="DNA")
```



Modeling Rate Variation Across Sites Proportion of Invariable Sites

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Modeling Rate Variation Across Sites Proportion of Invariable Sites

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p_inv ~ dnBeta(alpha_p, beta_p)

seq ~ dnPhyloCTMC(tree=psi, Q=Q, plnv=p_inv, type="DNA")

Can also add + I to any "standard" model of sequence evolution.



For scalar (numerical) parameters, we often report:

Mean

Median

95% Highest Posterior Density Interval



For trees, we often report:

Maximum A Posteriori (MAP) Tree ("Best" Tree)

mapTree(run1Trees)

For trees, we often report:

Maximum A Posteriori (MAP) Tree ("Best" Tree)

Majority-Rule Consensus Tree (Formed from all bipartitions with posterior > 0.5)

consensusTree(run1Trees,0.5)

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(Formed from all bipartitions with posterior > 0.5)

Greedy Consensus Tree

(Formed by ranking bipartitions and adding them until tree is fully resolved)

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Maximum A Posteriori (MAP) Tree ("Best" Tree)

Majority-Rule Consensus Tree

(Formed from all bipartitions with posterior > 0.5)

Greedy Consensus Tree

(Formed by ranking bipartitions and adding them until tree is fully resolved)

Maximum Clade Credibility Tree

(Sampled tree whose product of bipartition probabilities is greatest)

mccTree(run1Trees)

Assessing Topological Convergence



Assessing Topological Convergence



Visualizations of Tree Space

run1Trees = readTrees("run1.trees")
run2Trees = readTrees("run2.trees")
maxdiff([run1Trees,run2Trees])

Finds the biggest difference in estimated marginal posterior probabilities for all bipartitions across different replicates.





Tree 10



RevBayes Tutorial Setup

After logging on to your VM...

cd ~

```
cp -r moledata/revbayes/ ./
```

cd revbayes/genetree/

```
mv ./scripts/*.Rev .
```

RevBayes CTMC Tutorial

https://revbayes.github.io/tutorials/ctmc/

I strongly encourage you to try building at least one Rev script yourself for this tutorial, rather than simply running those that are already available.

Also note that in the scripts that are already written, the default number of replicate runs is 4. This is great for assessing convergence, but will cause analyses to run fairly slowly. You can reduce this to 2 to make sure that runs finish in a reasonable amount of time.

You'll also want to change the printgen option of mnModel and mnFile to 10 (rather than 1). Otherwise, output files get huge!

RevBayes CTMC Tutorial

https://revbayes.github.io/tutorials/ctmc/

Look at Table 1 in the tutorial above.

Using the model descriptions in the table, use the same principles we've been practicing to set up each of these models in RevBayes.

Fill out Table 2 for as many models as you can.

rb mcmc_JC.Rev

End