## The Scientist Game

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- 1956: Eleusis invented by Robert Abbott
- 1977: Described by Martin Gardner Scientific American
- c. 1993: I adapted Scientist Game for enrichment activities in my daughters' elementary school classes
- 1994+: I started seeing how various collaborators fared

   Statistician, oncologist, physicist, biologist colleagues
   Graduate students in biostatistics, public health



- A simplified universe
  - One dimensional universe observed over time
  - Each position in the universe has an object
  - Goal is to discover any rules that might determine which objects are in a given location at a particular time



 Objects in the universe have only three characteristics, each with only two levels

- Color: White or Orange
- Size: BIG or small
- Letter: A or B

So only 8 kinds of objects in the universe:

A a B b A a B b

## **Universal Laws**

 The level of each characteristic (color, size, letter) for the object at any position in the universe is either

> completely determined by the prior sequence of that characteristic for objects at that position, OR

• is completely random (anything is permissible)

(No patterns involving probabilities less than 1)

(Adjacent positions have no effect)

## **Universal Laws**

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 Furthermore any pattern to the objects at a position over time is "stationary"

 The exact pattern repeats itself over a finite period of time (the "cycle")

 The following "pattern" is not considered possible, because the exact same sequence does not re-appear

Color only (a cycle of length 2):

## b a A b a a a B A a A A

The next object in the sequence must be white, but any size or letter will do:

a A b B

• Size and letter (a cycle of length 4):

## A a B b A a B b A a B b

The next object in the sequence must be a big A, but any color will do

A A

Size only (a cycle of length 2):

## B a B b A b B a B b A a

The next object in the sequence must be big, but any color or letter will do:

A B A B

No discernible pattern (in available data):

### A a b a A b B a B B A a

 If there is truly no deterministic pattern, then any object may appear next:

a A b B a A b B



- Goal is therefore to decide for some position
  - whether a rule governs the level of each characteristic, and

- if so, what that rule is (pattern to the sequence)

## **Hypothesis Generation**

Initially we have observational data gathered over time

 Amount of available information varies from position to position

 We want to identify some position that is the most likely to be governed by some deterministic rule

# **Observational Data**

Time
Pstn -11 -10 -9 -8 -7 -6 -5 -4 -3 -2 -1 0 1 2

114.	b	a	A	6	a	a	B	a	A	A	a	b	?	?
115.	A	a	B	b	A	a	۵	b	A	a	a	A	?	?
116.							B	b	A	a	a	6	?	?
117.						b	B	A	b	6	6	a	?	?
118.	A	b	B	A	B	b	A	B	B	a	B	B	?	?
119.		B	b	B	b	A	a	A	a	B	A	b	?	?
120.										B	B	6	?	?
121.				B	A	a	B	b	b	a	6	A	?	?

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## **Observational Data**

#### Time Pstn -11 -10 -9 -8 -7 -6 -5 -4 -3 -2 -1 0 1 2

....

.....

#### 118. A b B A B b A B B a B B ? ?



- Further observation?
  - Might take too long
  - Won't really establish cause and effect

## Experimentation

You can try to put an object in the position

- If it cannot come next, it disintegrates and you can try another
- If it can come next, it stays and you can try a different object to follow it

- Ultimately, a sequence of experiments can be used



You need to devise a series of experiments to discover

 whether a deterministic rule governs the sequence of objects at position 118, and

- if there is such a rule, what it is



Problem:

You must buy objects to experiment with

- (apply for a grant)
- Question:

– What object should you try next in the sequence in order to determine the rule?

## **Possible Experiments**

Time Pstn -11 -10 -9 -8 -7 -6 -5 -4 -3 -2 -1 0 1 2 118. A b B A B b A B B a B P ? ?

## <u>Possible Experiments</u> a A b B a A b B

Which experiment do you do first?

## **Reviewing the Grant Application**

- Did you choose a good experiment?
- In order to determine whether your grant application should be funded, we review an ideal scientific approach

1. Observation

- 2. Formulating hypotheses
- 3. Devising experiments which discriminate between hypotheses

## **Results of Observation**

 Time

 Pstn -11 -10 -9 -8 -7 -6 -5 -4 -3 -2 -1 0 1 2

 118.
 A
 b
 B
 A
 B
 B
 B
 B
 C
 P

- We identified position 118 which had some regular patterns
  - Color cycle of length 2: (orange, white)
  - Size cycle of length 4: (big, little, big, big)

Letter cycle of length 3: (A, B, B)

# **Define Hypotheses**

Deterministic pattern vs random chance

 Some (or all) of the observed patterns for each characteristic might be coincidence

- Is coincidence realistic?
  - Assume each level equally likely and a sample size of 12
  - Chance of observing a pattern for a single characteristic
    - 1 out of 1,024 for a cycle of length 2
    - 1 out of 512 for a cycle of length 3
    - 1 out of 256 for a cycle of length 4
    - 1 out of 134,217,728 for all three simultaneously

## **Possible Hypotheses**

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Assuming sufficient data to see any rule
 <u>118.</u> A b B A B b A B B a B B ? ?

Hypotheses Color, Size, and Letter Color, Size Color, Letter Size, Letter Color Size Letter All coincidence

### 118. A b B A B b A B B a B B ? ? Possible Experiments Hypotheses A Color, Size, and Letter Color, Size Color, Letter Size, Letter Color Size Letter All coincidence

### 118. A b B A B b A B B a B B ? ? Possible Experiments Hypotheses A Color, Size, and Letter + Color, Size Color, Letter Size, Letter Color Size Letter All coincidence

<u>118.</u>	A	b	B	A	B	b	A	B	B	a	B	B	?	?
							<u>P</u>	<u> 055</u>	ib]	<u>e E</u>	<u>xpe</u>	rim	<u>ent</u>	<u>s</u>
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<u>118.</u>	A	b	B	A	B	b	A	B	B	a	B	B	?	?
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<u>118.</u>	A	b	B	A	B	b	A	B	B	a	B	B	?	?
							<u>P</u>	<u> 055</u>	ib]	<u>e E</u>	xpe	<u>r in</u>	<u>ent</u>	<u>S</u>
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Color,	Lett	er					+							
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 A noninformative experiment **118.** A b B A B b A B B a B B ? ? Possible Experiments Hypotheses A Color, Size, and Letter + Color, Size + Color, Letter + Size, Letter +Color +Size + Letter + All coincidence +

## **Next Worse Choice**

 If all hypotheses equally likely, a 7-1 split **b B A B b A B b a B** 118. A B ? Possible Experiments A b Hypotheses Color, Size, and Letter + -Color, Size \_ + 35 Color, Letter + \_ Size, Letter + -Color + Size + -Letter All coincidence

# If Eliminate Hypotheses 1 by 1

Guessing a number between 1 and 1,000

You can ask Yes-or-No questions

Strategy 1: Elimination 1 by 1

- "Is it 137? (NO) Is it 892? (NO) ..."
- On average it will take 500 questions

#### Strategy 2: Binary search

- Split remaining hypotheses in half each time
- "Is it > 500? (NO) Is it > 250? (YES) Is it > 375?..."
- You can know the answer after 10 questions (2<sup>10</sup>=1,024)

## **Other Suboptimal Experiments**

 If all hypotheses equally likely, a 6-2 split **b B A B b** A B B a B 118. B ? A Possible Experiments Hypotheses b b B a A Color, Size, and Letter -Color, Size Color, Letter Size, Letter Color Size Letter

All coincidence

## **Optimal Experiments**

 Based on a binary search 6 b B B 118. B B A B A A a B Possible Experiments Hypotheses b 0 B a B a A A Color, Size, and Letter Color, Size Color, Letter -Size, Letter +Color + -Size + Letter +All coincidence

## Interpreting Good Experiments

 We can easily describe what we were testing for in the three "best" experiments

Is Letter important?
We used the size and color that would work regardless

- Is Size important?
  - We used the letter and color that would work regardless
- Is Color important?
  - We used the size and letter that would work regardless

## Sequence of Experiments

- Separate question into three experiments
  - Address each characteristic separately
  - Avoid "confounding" the question
- Perform these 3 experiments in sequence

   Results uniquely identify the 8 hypotheses
   (Eliminating hypotheses 1 at a time would on average take 4 experiments)
- (No other series of 3 will always do this)

## **Optimal Experiments**

 Based on a binary search 118. A b B A B b A B B a B B ? ? Possible Experiments **Hypotheses** B a A Color, Size, and Letter Color, Size -Color, Letter + Size, Letter -+Color + + -Size + - +Letter + + All coincidence

## **Other Experimental Sequences**

 Based on results from first experiment, additional good experiments for the second stage possible, e.g.,

Suppose first experiment: A <u>and</u> it does not disintegrate

- Then we know color does not matter
- Good choices for the next experiment: **B a B a** 
  - Choose different letter or size, but not both
- BUT: If first experiment had disintegrated, only two good choices: B a
  - Must use orange, because color matters

## Supposing A Works

 Based on a binary search 118. A b B A B b A B B a B B Possible Experiments **Hypotheses** B a B a A Color, Size, and Letter Color, Size Color, Letter Size, Letter Color Size Letter All coincidence

## **Bayesian Considerations**

- With a binary search, we want to eliminate 50% of the hypotheses based on their "prior probability"
- If we had a prior belief that most of our universe was random chance, then perhaps b might be a good first experiment
- We can cook up an example to try to make that a good Bayesian choice when we have seen (perhaps spurious) patterns like 118

## Toy Example

Assume 99.85% of positions truly have no pattern

•

- Others have "independent, equiprobable patterns" with cycle length < 12</li>
- Expect to see 0.00000002% with patterns like 118
   Maybe we examined millions of positions
- Best approach may have been to try:
  - Discriminates between no pattern (like 49% of positions) and some pattern (like the other 51% of positions)
  - On average, 2.52 experiments (1 expt 49%, 4 expt 51%)

# **Details of Toy Example**

Not for the faint of heart, but...

Suppose color, letter, size independent

For each factor

- 99.5% of sites have no pattern
- Rest equally likely to have cycles of 2, 3, or 4
- For every length of cycle, all patterns equally likely
  - E.g., for big white letters
    - Cycle length 2: AA, AB, BB each 1/3
    - Cycle length 3: AAB, ABB each 1/2
    - Cycle length 4: AAAB, AABB, ABBB each 1/3

## **Posterior Probabilities for 118**

 We chose a site with observed patterns of color cycle=2, letter cycle=3, size cycle=4

3.8%

3.8%

2.6%

0.9%

Of all such patterns, the truth will be

- All coincidence 49.5%
- Letter only 16.9%
- Size only 11.3%
- Color only 11.3%
- Letter, size only
- Color, letter only
- Color, size only
- Color, letter, size

## What If Data Insufficient?

 Suppose deterministic cycle length > 12 **b** 118. B A B B A 0 B B B a A Possible Experiments Hypotheses 6 6 B a B A a A Color, Size, and Letter Color, Size Color, Letter -Size, Letter + Color + -Size + Letter + All coincidence + ? ? ? 2 ? 2 Cycle length > 12 2

# If Cycle Length > 12

- We would have no information to be able to guess the true pattern
- BUT, we might gain some information from A as a first experiment (and thus partial vindication)
  - If A disintegrated we would know that there was some deterministic pattern with cycle length > 12
    - But we would have no clue about the pattern
  - And we have to question the efficiency of this strategy
    - Deterministic patterns of length > 12 might demand A and thus we cannot presume one of the other hypotheses is established

# Moral: Hypotheses

 The goal of the experiment should be to "decide which" not "prove that"

- A well designed experiment discriminates between hypotheses
  - The hypotheses should be the most important, viable hypotheses

# Moral: Experiment

 All other things being equal, an experiment should be equally informative for all possible outcomes

In the presence of a binary outcome, use a binary search

 (using prior probability of being true)

But need to consider simplicity of experiments, time, cost
 – (What lessons can be learned from Master Mind?)

## In the Presence of Variability

- We use statistics to quantify the precision of our inference
- We will describe our confidence/belief in our conclusions using frequentist or Bayesian probability statements
- Discriminating between hypotheses will be based on a frequentist confidence interval or a Bayesian credible interval

## **Interval Estimates**

Frequentist confidence intervals

- The set of all hypotheses for which the observed data are "typical"
- There is more than a negligible probability of obtaining such results when those hypotheses are true

### Bayesian credible intervals

- The set of hypotheses that are most probable given the observed data
- Must incorporate our prior belief in the hypotheses

## **Frequentist Evidence**

Does frequentist evidence provide evidence?

- Is it relevant to calculate the probability of data that you know you observed?
- Relevance especially questionable if calculated on a hypothesis that is unlikely a priori
- My answer in experimental design: Yes
  - Design an experiment that has results that are not consistent with one of the viable, important hypotheses

## Statistical Experimental Design

I believe a scientific approach to the use of statistics is to

 Decide a level of confidence used to construct frequentist confidence intervals or Bayesian credible intervals

 Ensure adequate statistical precision (sample size) to discriminate between relevant scientific hypotheses

 The intervals should not contain two hypotheses that were to be discriminated between

## Impact on Statistical Power

 In evaluating a design, I always examine the alternative hypothesis for which we have I equal one-sided type I and type II errors

- E.g., 97.5% power to detect the alternative in a one-sided level 0.025 hypothesis test
- In this way, at the end of the study, the 95% CI will not contain both the null and alternative hypotheses
  - I will have discriminated between the hypotheses with high confidence