## A European application of random matrix theory

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## Genetics

[Novembre et al., Nature 2008]: Researchers analyzed genetic data from people with European ancestry: 1,387 people at 197,146 genetic loci.



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They then analyzed this data by examining the singular values of X....

# **Probability Theory**

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## **Unfair Coins:**



https://izbicki.me/blog/how-to-create-an-unfair-coin-and-prove-it-with-math.html

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Probability theory: Provides the tools needed to compute the expected outcome.

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## **Random Matrix Theory**

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$$A = \left(\begin{array}{cc} x & 0 \\ 0 & y \end{array}\right) \qquad \qquad B = \left(\begin{array}{cc} 1 & y \\ y & 1 \end{array}\right)$$

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Since the matrix is (at least partially) random, the data associated to the matrix is potentially random as well: the entries of X, det(X), Tr(X), eigenvalues of X, eigenvectors of X, etc.

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$$(0, -1),$$
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Can still find the least squares solution using this random matrix:

$$\left(\begin{array}{cc} 4+q^2 & 2+q\\ 2+q & 3 \end{array}\right)\left(\begin{array}{c} a\\ b \end{array}\right) = \left(\begin{array}{c} 6+2q\\ 4 \end{array}\right)$$

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## Least Squares? (continued)









McConnell

Not McConnell





Eugene Wigner

Sad McConnell

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- Wigner's idea: treat the Hamiltonian as a random matrix

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n = 10,000

### Semicircle Distribution

The histograms get closer and closer to the semicircle distribution:



### Theorem (Wigner's Semicircle Law)

Let  $X_n$ ,  $n \in \mathbb{N}$ , be the sequence of symmetric random matrices as above. For any interval  $[a, b] \subset \mathbb{R}$ ,

$$\lim_{n \to \infty} \frac{\#\{\text{eigenvalues of } X_n \text{ in the interval } [a, b]\}}{n} = \int_a^b s(t) \ dt.$$

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That is, there exists an  $N \in \mathbb{N}$  so that for any  $n \geq N$ 

$$\frac{\#\{\text{eigenvalues of } X_n \text{ in the interval } [a,b]\}}{n} \approx \frac{1}{2\pi} \int_a^b \sqrt{4-t^2} \ dt.$$

## The Marčenko–Pastur Law



Vladimir Marčenko



Leonid Pastur

#### Marčenko–Pastur distribution

Fix  $\lambda \in (0,1]$  and define  $\lambda_{\pm} = (1 \pm \sqrt{\lambda})^2$ . The Marčenko–Pastur distribution is

$$u(t) := egin{cases} rac{1}{2\pi\lambda}rac{\sqrt{(t-\lambda_-)(\lambda_+-t)}}{t} & ext{if } \lambda_- \leq t \leq \lambda_+ \ 0 & ext{otherwise} \end{cases}$$



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#### Theorem (The Marčenko–Pastur Law)

With  $Y_n$ ,  $n \in \mathbb{N}$ , as above, for any interval  $[a, b] \subset \mathbb{R}$ ,

$$\lim_{n\to\infty}\frac{\#\{\text{eigenvalues of }Y_n \text{ in the interval }[a,b]\}}{p(n)} = \int_a^b \nu(t) \ dt.$$

That is, there exists an  $N \in \mathbb{N}$  so that for any  $n \geq N$ 

$$\frac{\#\{\text{eigenvalues of } Y_n \text{ in the interval } [a,b]\}}{p(n)} \approx \frac{1}{2\pi\lambda} \int_a^b \frac{\sqrt{(t-\lambda_-)(\lambda_+-t)}}{t} \ dt.$$

## **Back to Genetics**

Researchers recorded genetic data as the matrix

	locus 1	locus 2	· · · I	ocus j	•••	locus 197146	
person 1	$\int x_{1,1}$	<i>x</i> <sub>1,2</sub>	• • •		•••	<i>X</i> 1,197146	)
person 2	<i>x</i> <sub>2,1</sub>	<i>x</i> <sub>2,2</sub>				÷	
÷	:	÷	·				=: X
person i				x <sub>i,j</sub>			
÷	:				·	÷	
person 1387	$(x_{1387,1})$	•••			• • •	<i>x</i> <sub>1387,197146</sub>	)

Researchers recorded genetic data as the matrix

Set

$$\begin{array}{l} n := 197, 146 \\ p(n) := 1,387 \end{array} \qquad \lambda := \frac{p(n)}{n} = \frac{1,387}{197,146} \approx 0.007035 \end{array}$$

They computed eigenvalues of  $Y := \frac{1}{n}XX^T$ , which is a  $p(n) \times p(n)$  symmetric matrix.

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- Something pretty incredible happens when you plot these coordinate pairs...



