

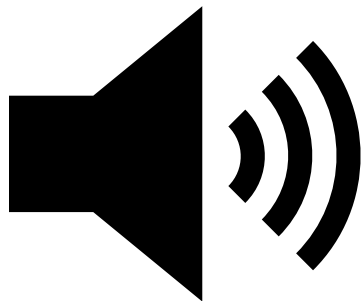
# See elegance in a worm brain



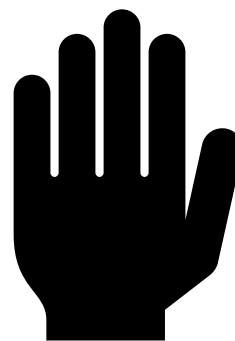
Designated assistant professor  
Graduate school of Science, Nagoya University

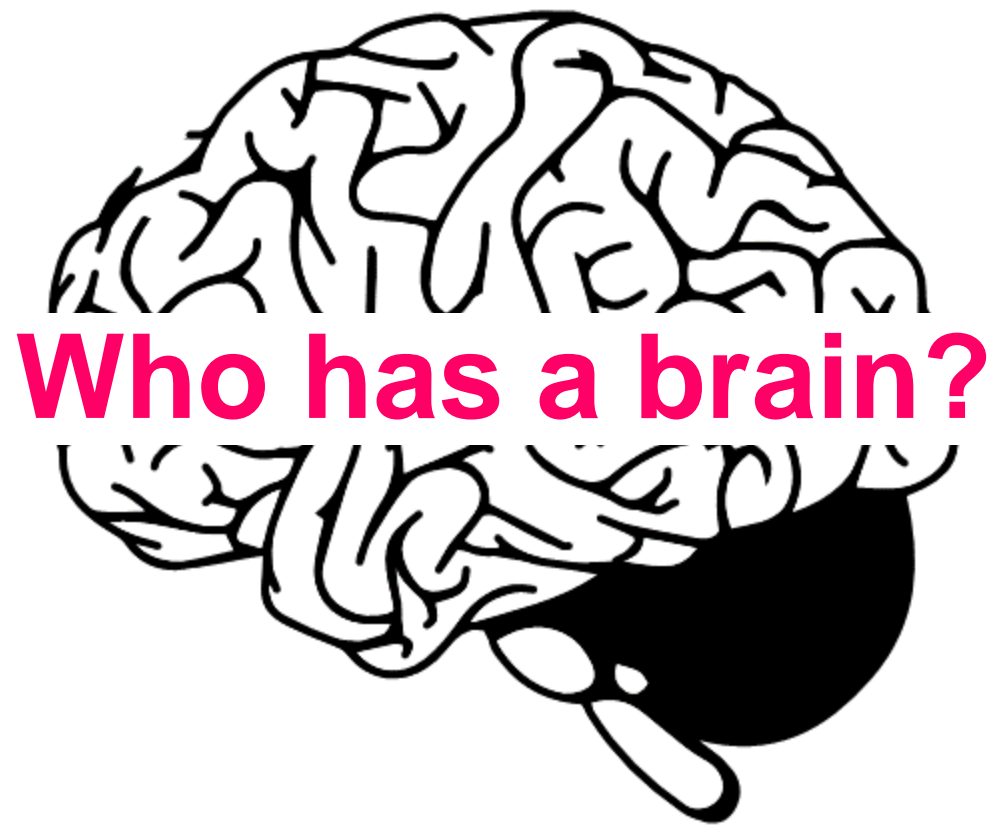
**Ken Noma**

**Volume**



**STOP!**





**Who has a brain?**

A black and white photograph of a starry night sky. In the upper left quadrant, there is a large, bright, hazy nebula or star cluster. The rest of the sky is filled with numerous stars of varying brightness, some appearing as sharp points of light and others as soft, out-of-focus spots. The overall tone is dark and grainy, typical of a long-exposure astronomical photograph.

**Wonderful!!**

# Today's menu

- *Appetizer* -  
**Brain**



- *Main dish* -  
**Worm**

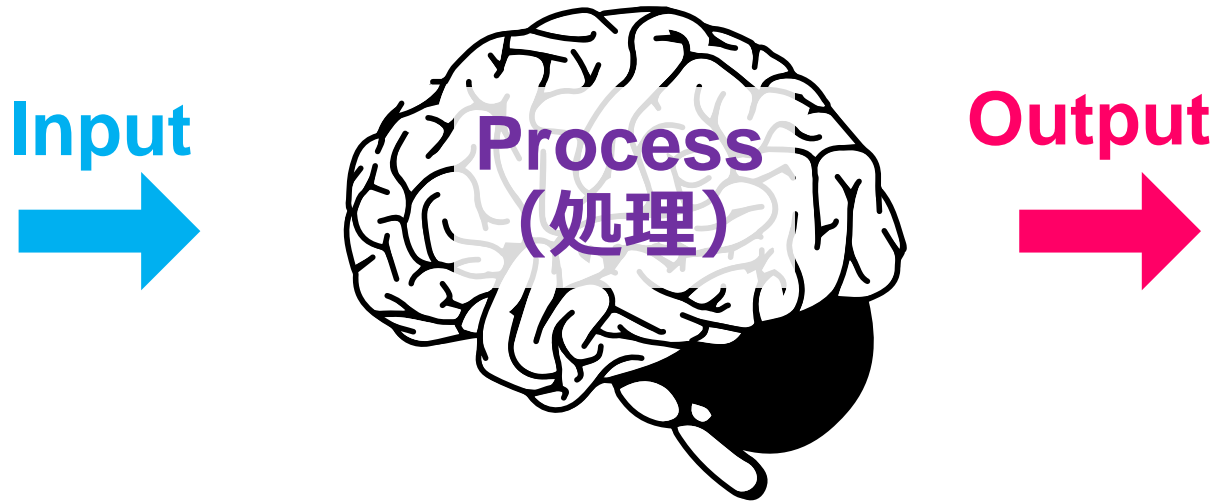


- *Dessert* -  
**Me**



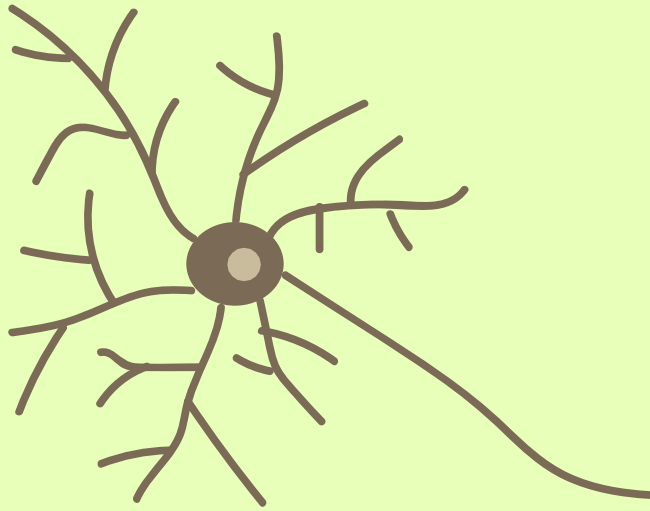
**8 elegant  
things**

# The brain is

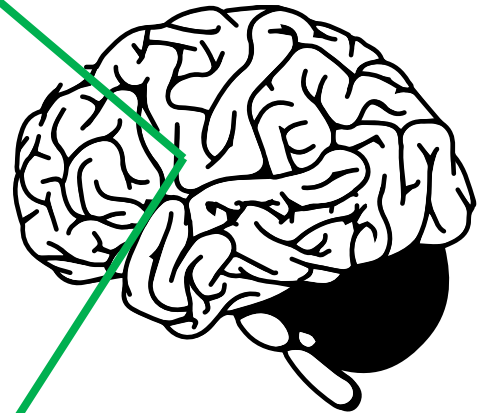


the organ inside the head that controls thought, memory, feelings, and activity. (Cambridge dictionary)

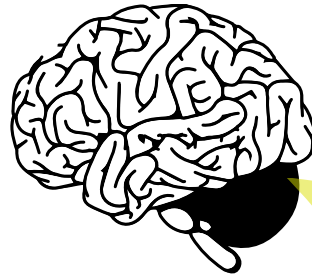
# Neurons are the functional units of the brain



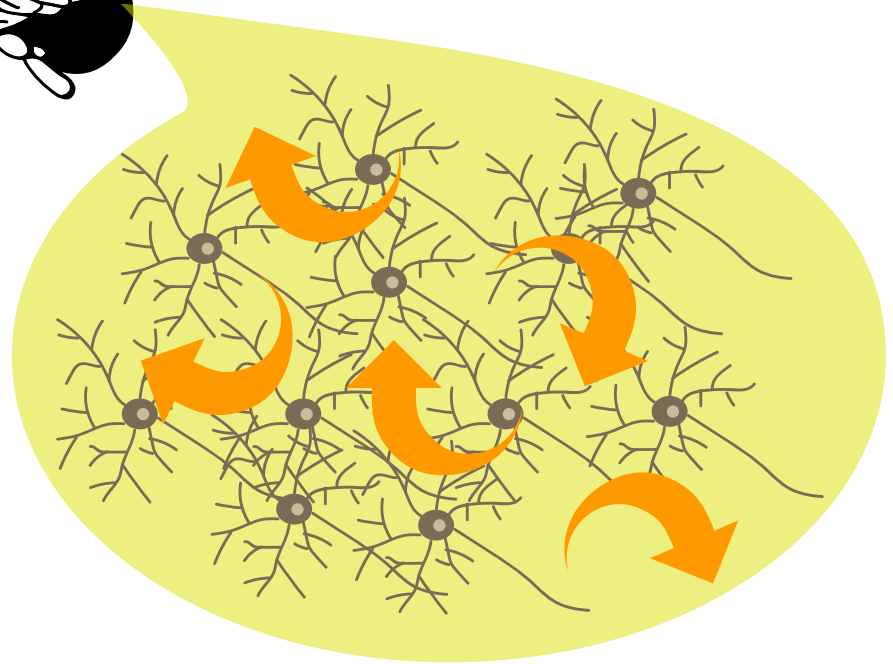
**Neuron 神經細胞**  
(a type of the cell)



# Neurons communicate with each other to process information



**Communication**





# Exciting questions about the brain

How do neurons process information?  
処理する

How is the brain formed?

What is the consciousness?  
意識



**The problem**

**Our brain is too complex**

# Quiz

## How many neurons in our brain?

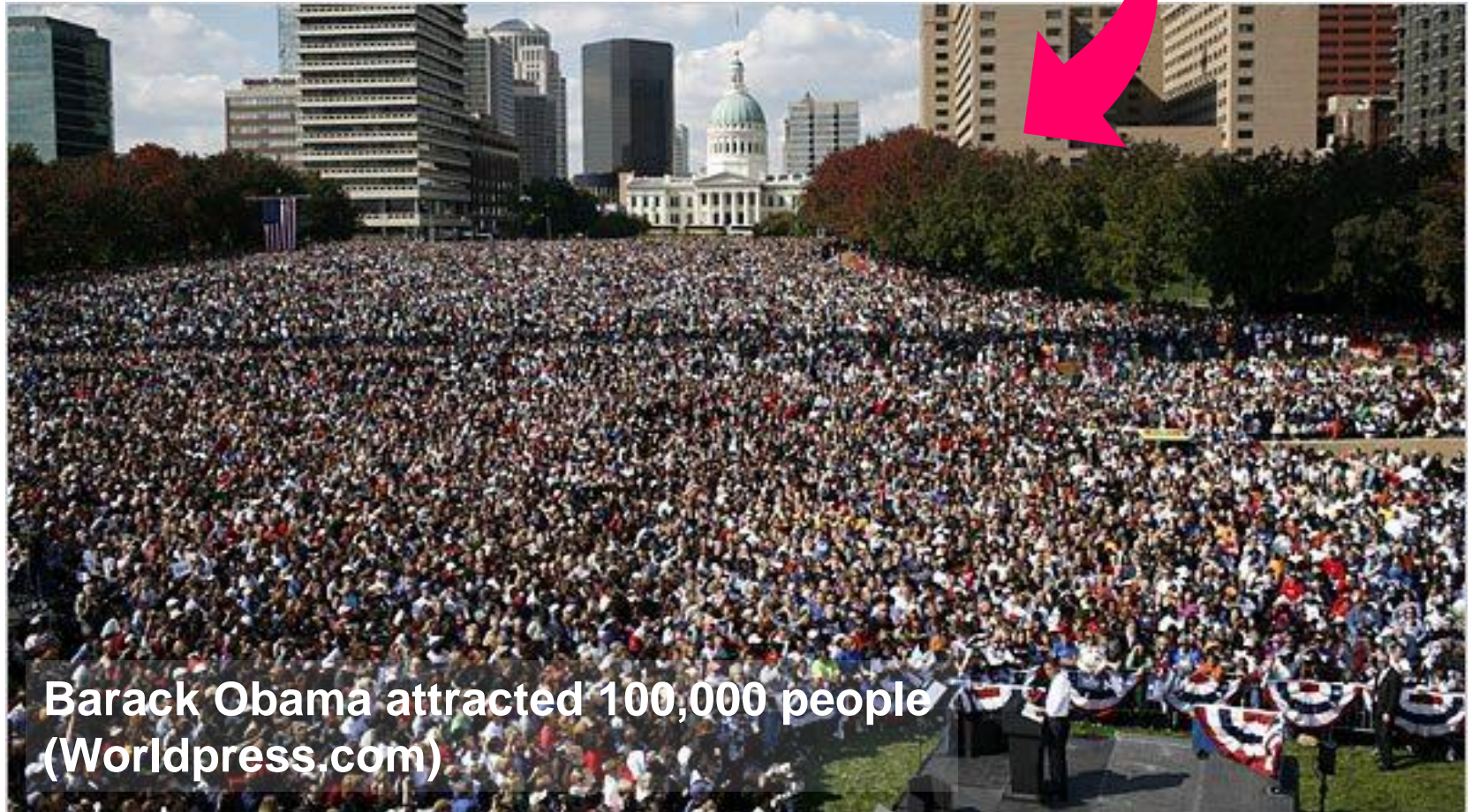
1. 1000 (thousand)
2. 1000,000 (million)
3. 1000,000,000 (billion)

# Quiz

## How many neurons in our brain?

1. 1000 (thousand)
2. 1000,000 (million)
3. 1000,000,000 (billion)
4. 100,000,000,000 (100 billion)

# 1,000,000 times this



Barack Obama attracted 100,000 people  
(Worldpress.com)



by Chip Somodevilla/Getty.

**That's...  
too...  
complicated...**

# Problems of the human brain as a research subject

Human brain is

1. too complicated



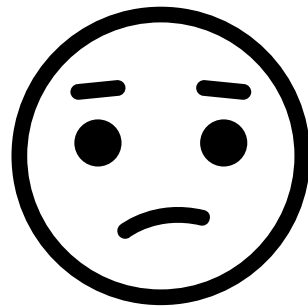
2. hard to see



3. hard to manipulate



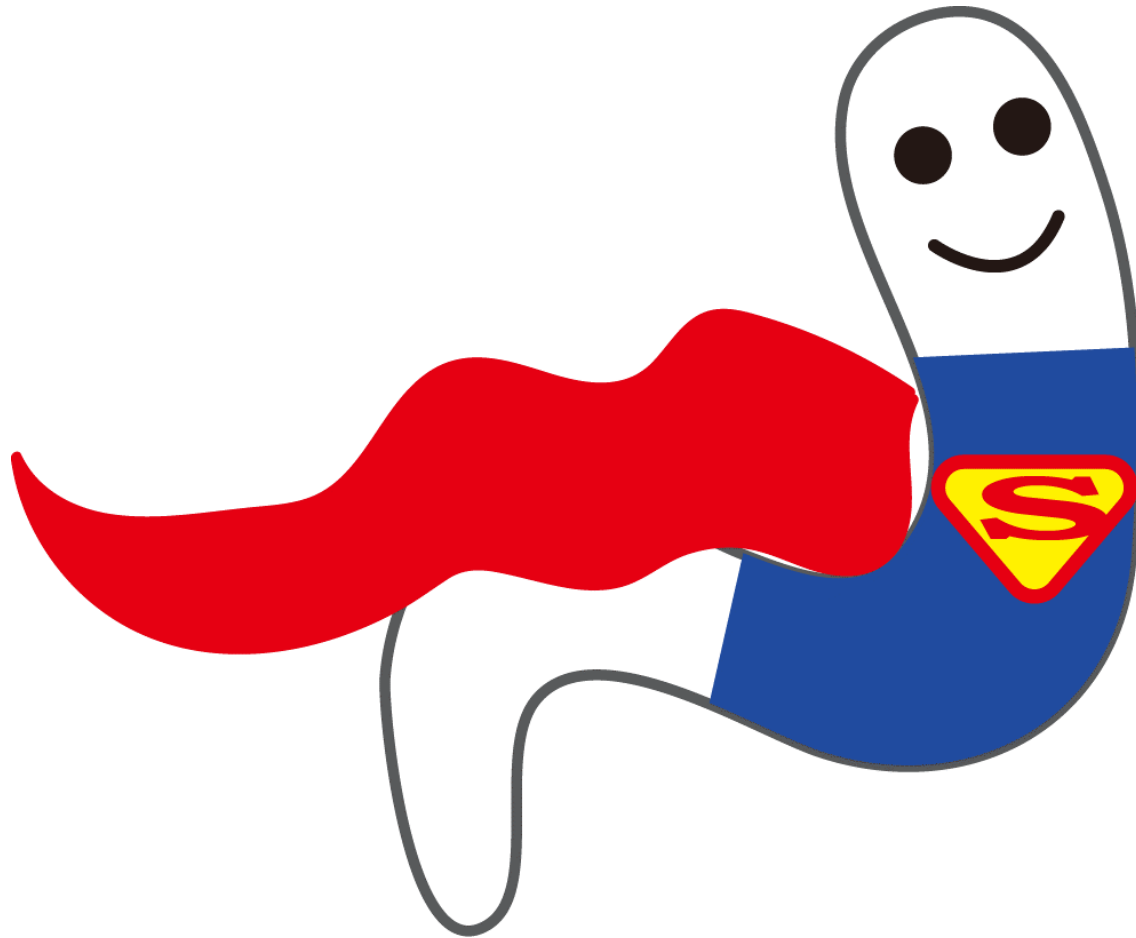
# What should we do?



We need  
a hero...



# Worm the hero



# Today's menu

- *Appetizer* -

**Brain**



- *Main dish* -

**Worm**



- *Dessert* -

**Me**

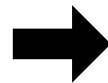


# The worm

## *Caenorhabditis elegans* (*C. elegans*)

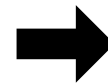
### Model organism

Simple and easy!!



Core principle

原理



# The worm

## *Caenorhabditis elegans* (*C. elegans*)



Sydney Brenner

1974

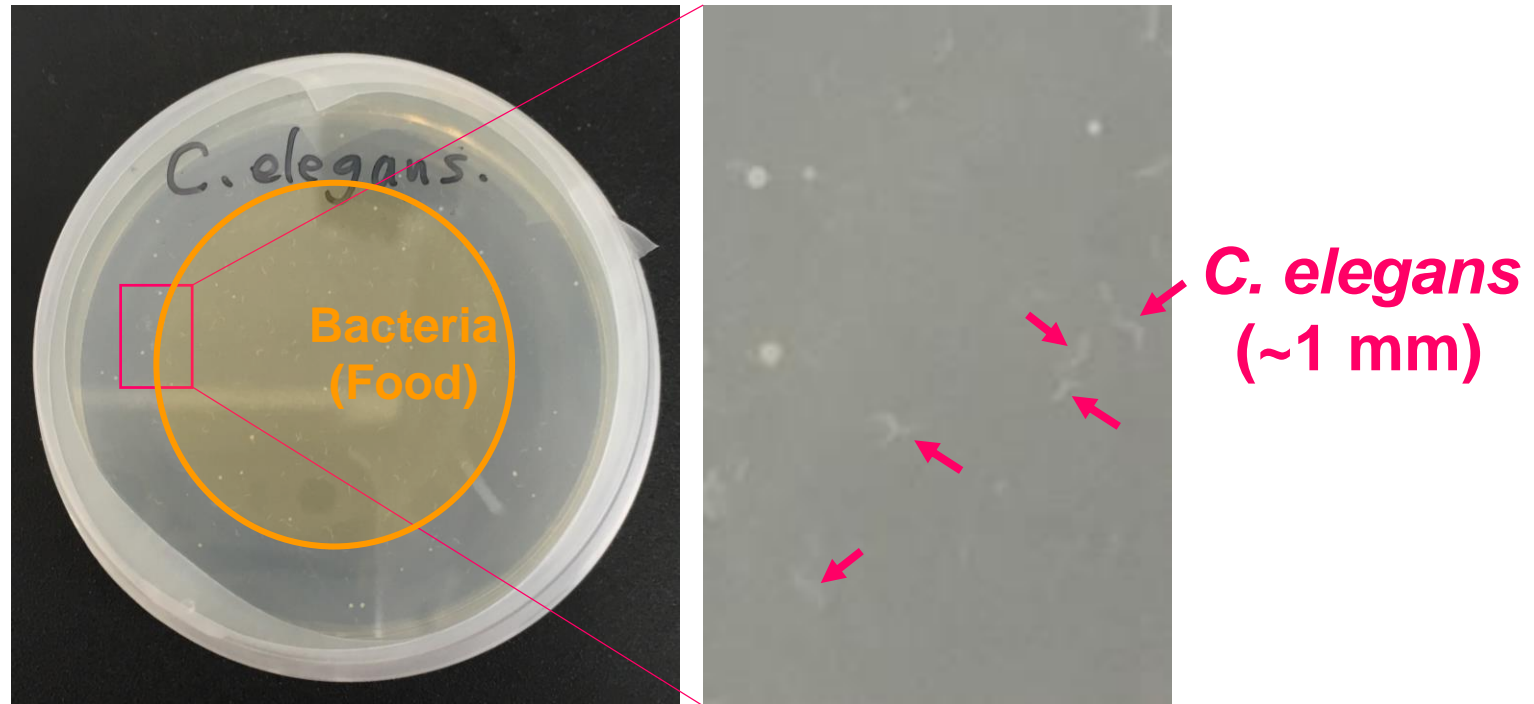
THE GENETICS OF *CAENORHABDITIS ELEGANS*

S. BRENNER

*Medical Research Council Laboratory of Molecular Biology,  
Hills Road, Cambridge, CB2 2QH, England*

**Nobel prize in 2002**

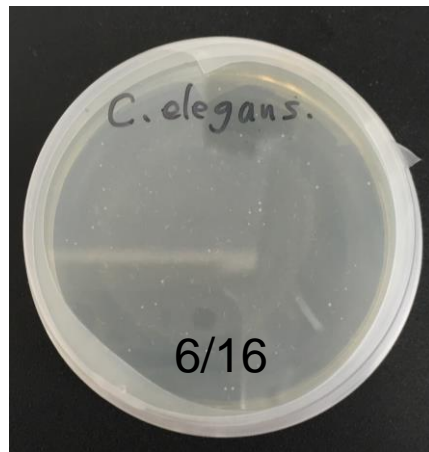
# You are going to receive this



*C. elegans* is **NOT** parasitic **NOR** harmful.  
寄生性

# Elegance 1

## *C. elegans* is easily maintained (飼育)



Hermaphrodite  
雌雄同体



3 days  
↓

~1000 worms



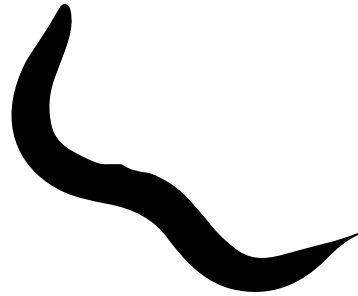
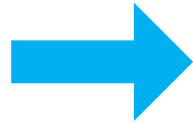
# The worm show



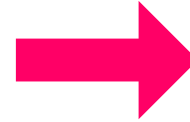
## Elegance 2

*C. elegans* senses and responds to stimuli  
(刺激)

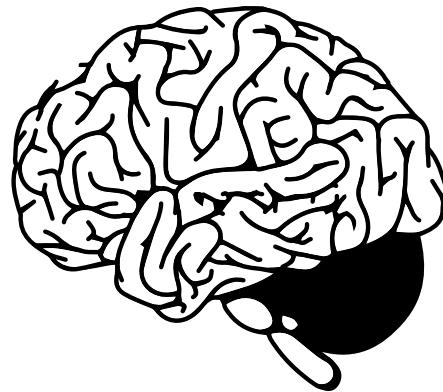
Touch



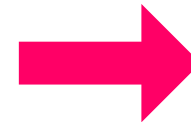
Movement



Input



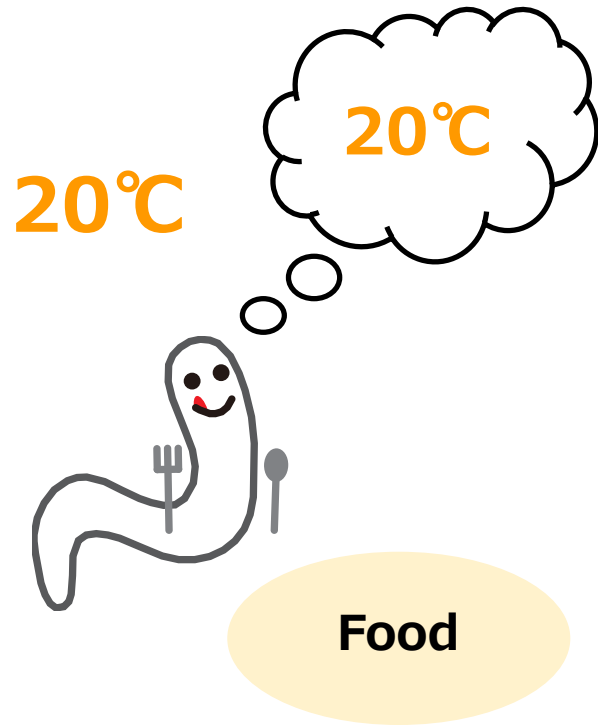
Output





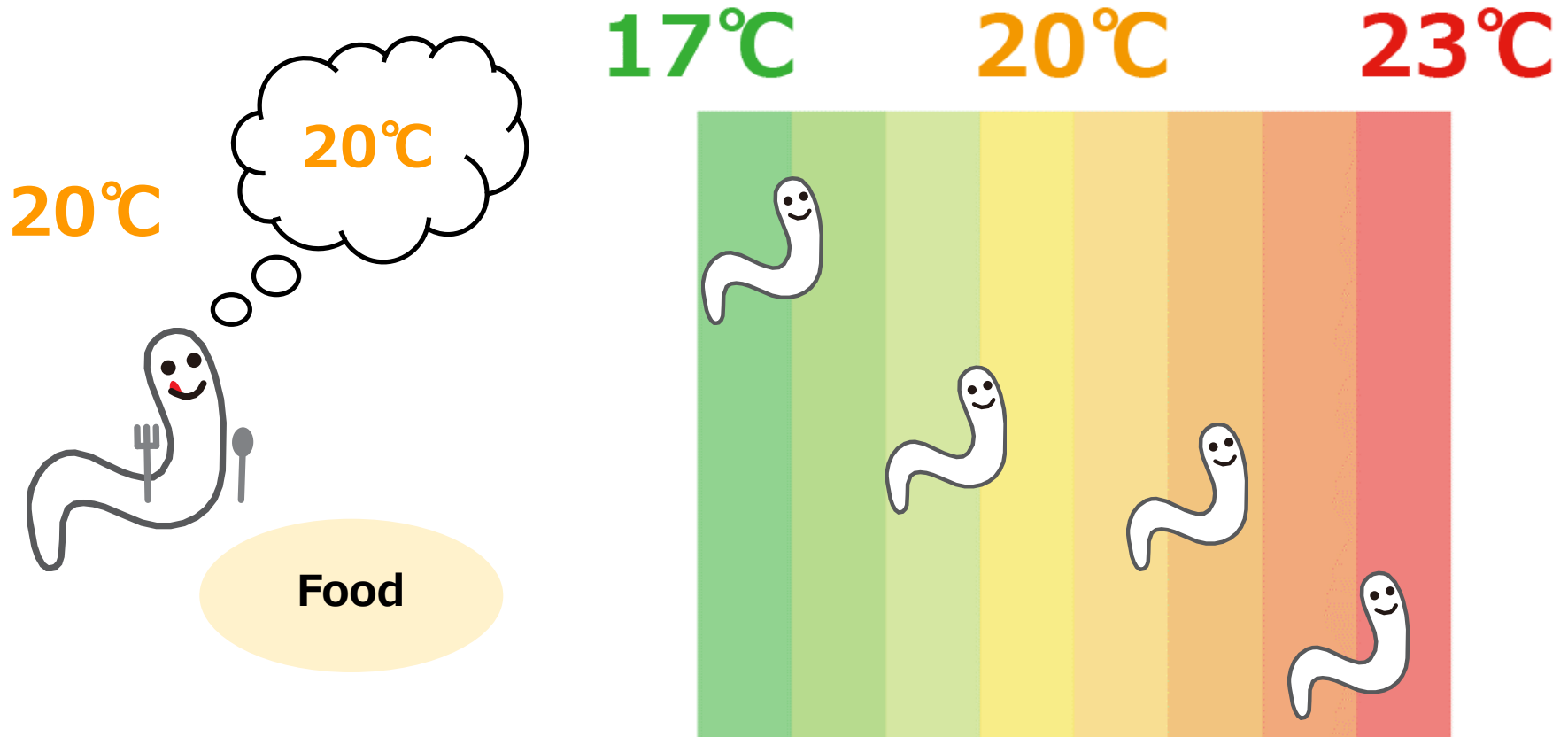
# Elegance 3

## *C. elegans* has memory



# Elegance 3

## *C. elegans* has memory



Hedgecock and Russell, *PNAS*, 1975  
Mori and Ohshima, *Nature*, 1995

# Elegance 4

## *C. elegans* has a brain



**Elegance 5**  
**Every *C. elegans* is the same**

**302** neurons



Good for experiments!!

**Well, *C. elegans* seems to be good for experiments, but...**

**Who cares?**

**What can we learn?**



# Exciting questions about the brain

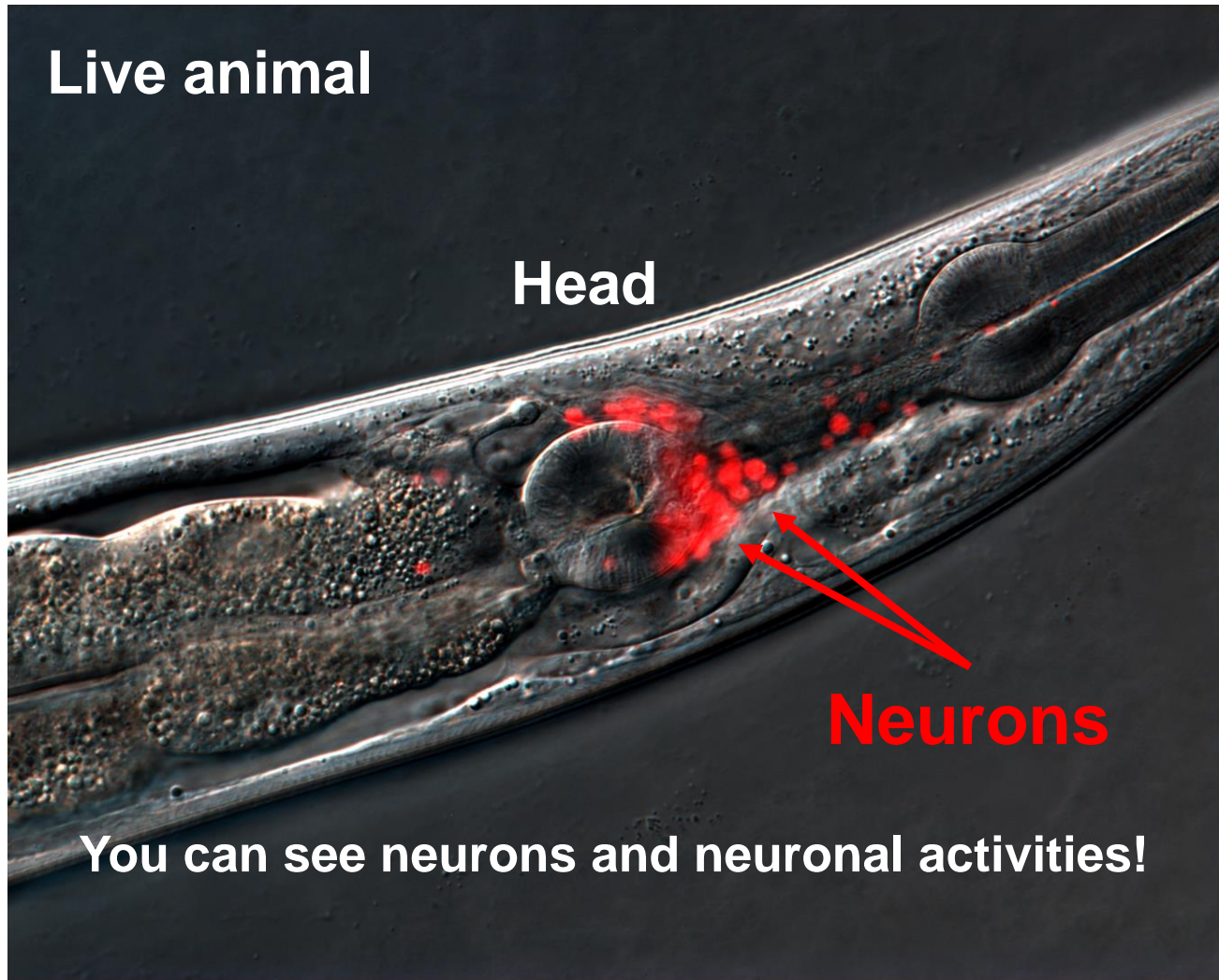
**How do neurons process information?**

How is the brain formed?

What is the consciousness?

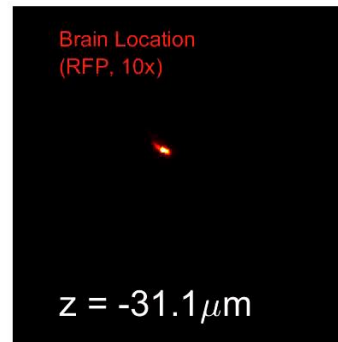
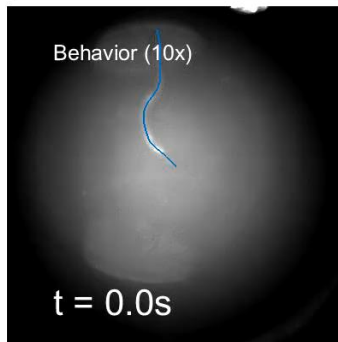
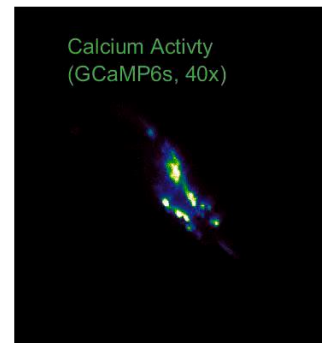
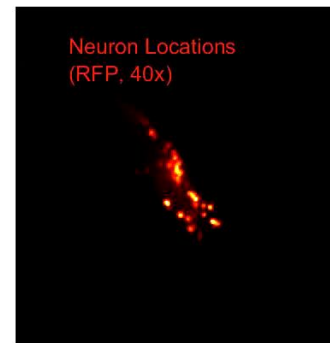
# Elegance 6

## *C. elegans* is transparent



# Elegance 6

Whole brain activities can be imaged in a freely moving animal



← Neuronal activity  
=  
What *C. elegans*  
is “thinking”.

Nguyen et al., *PNAS*, 2016

Neuronal activity



Core principle of  
information processing



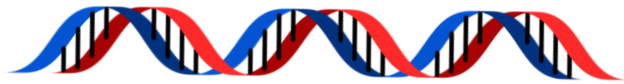
# Exciting questions about the brain

How do neurons process information?

**How is the brain formed?**

What is the consciousness?

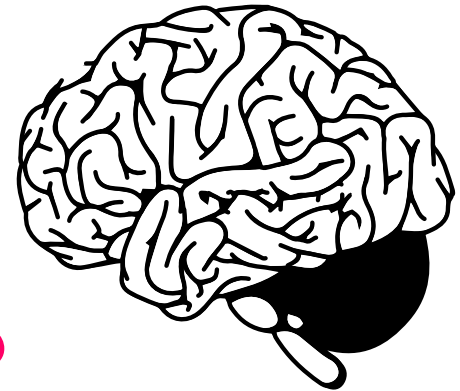
# The brain is formed, based on the genome



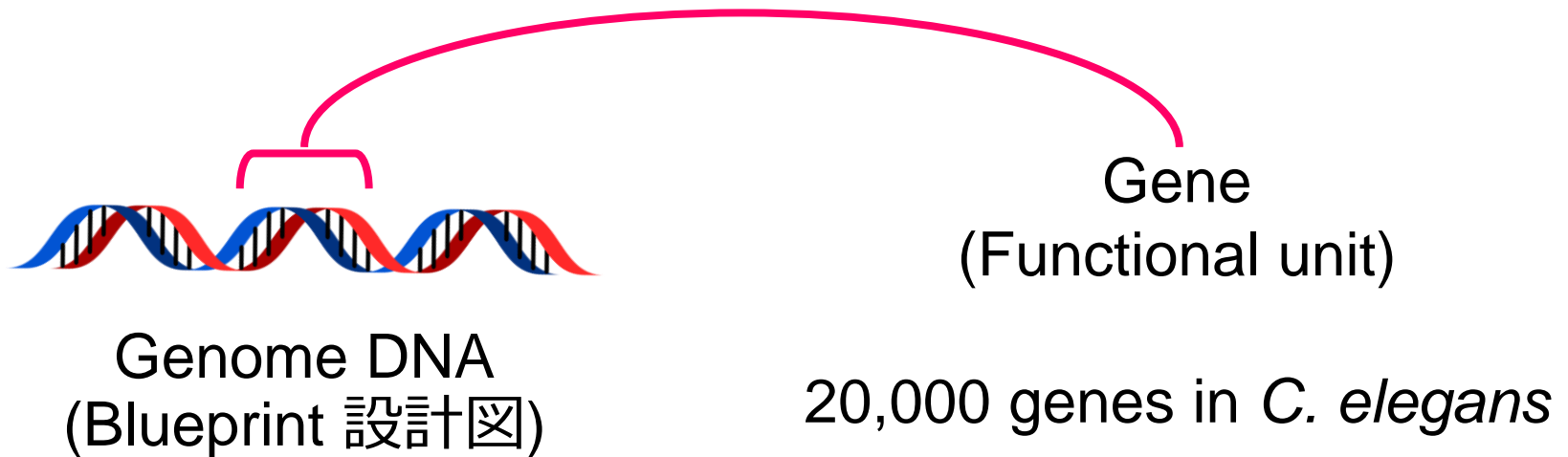
Genome DNA  
(Blueprint 設計図)



**HOW??**



# Genes are the functional units of the genome



Which gene is important for the brain??

# Elegance 7

***C. elegans* is easily manipulated**

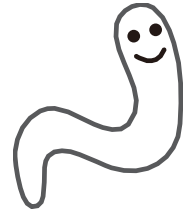
# Elegance 7

## *C. elegans* is easily manipulated

Blueprint  
設計図



Gene A

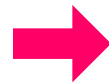


Gene B



Gene C

Forward genetic screen

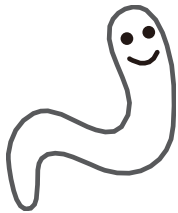


Important for the brain!

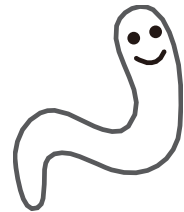
# Elegance 7

## *C. elegans* is easily manipulated

Blueprint  
設計図



Gene A

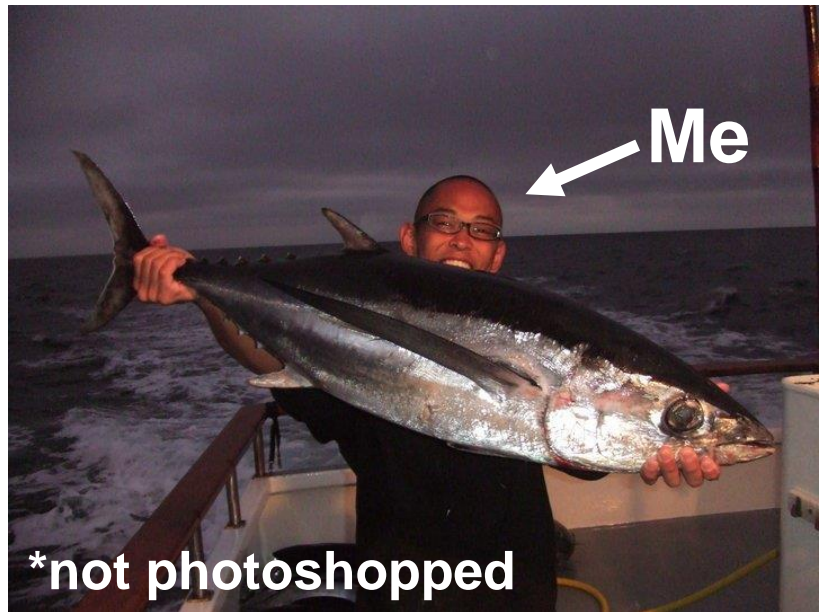


Gene B



Gene C

Forward genetic screen



Important for the brain!

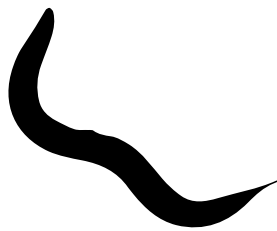
# Example: a gene, *unc-13* important for neuronal communication



# Elegance 8

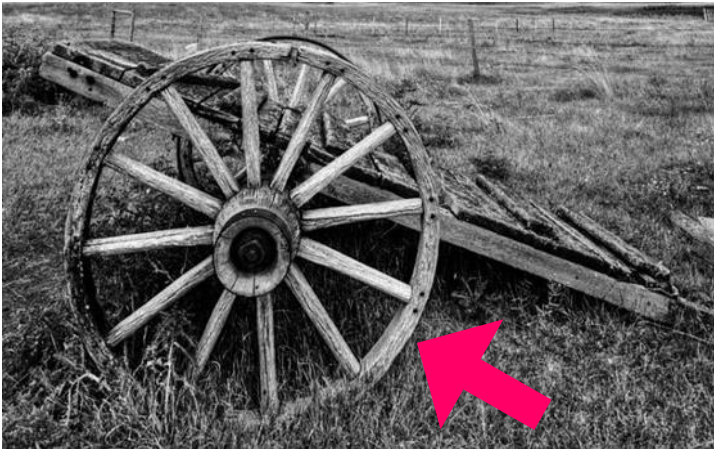
## Important genes are conserved

保存されている



The blueprint

***unc-13***



[https://books.google.co.jp/books/about/Twist\\_of\\_Gold.html?id=tJKbQis8-xEC&source=kp\\_cover&redir\\_esc=y](https://books.google.co.jp/books/about/Twist_of_Gold.html?id=tJKbQis8-xEC&source=kp_cover&redir_esc=y)  
2018/10/18

<https://response.jp/article/img/2018/02/23/306453/1279152.html?from=arrow-next> 2018/10/18



# Take-home message

**“Model organisms” are powerful!!**



**Choose a  
right tool!!**

# Today's menu

- *Appetizer* -

**Brain**



- *Main dish* -

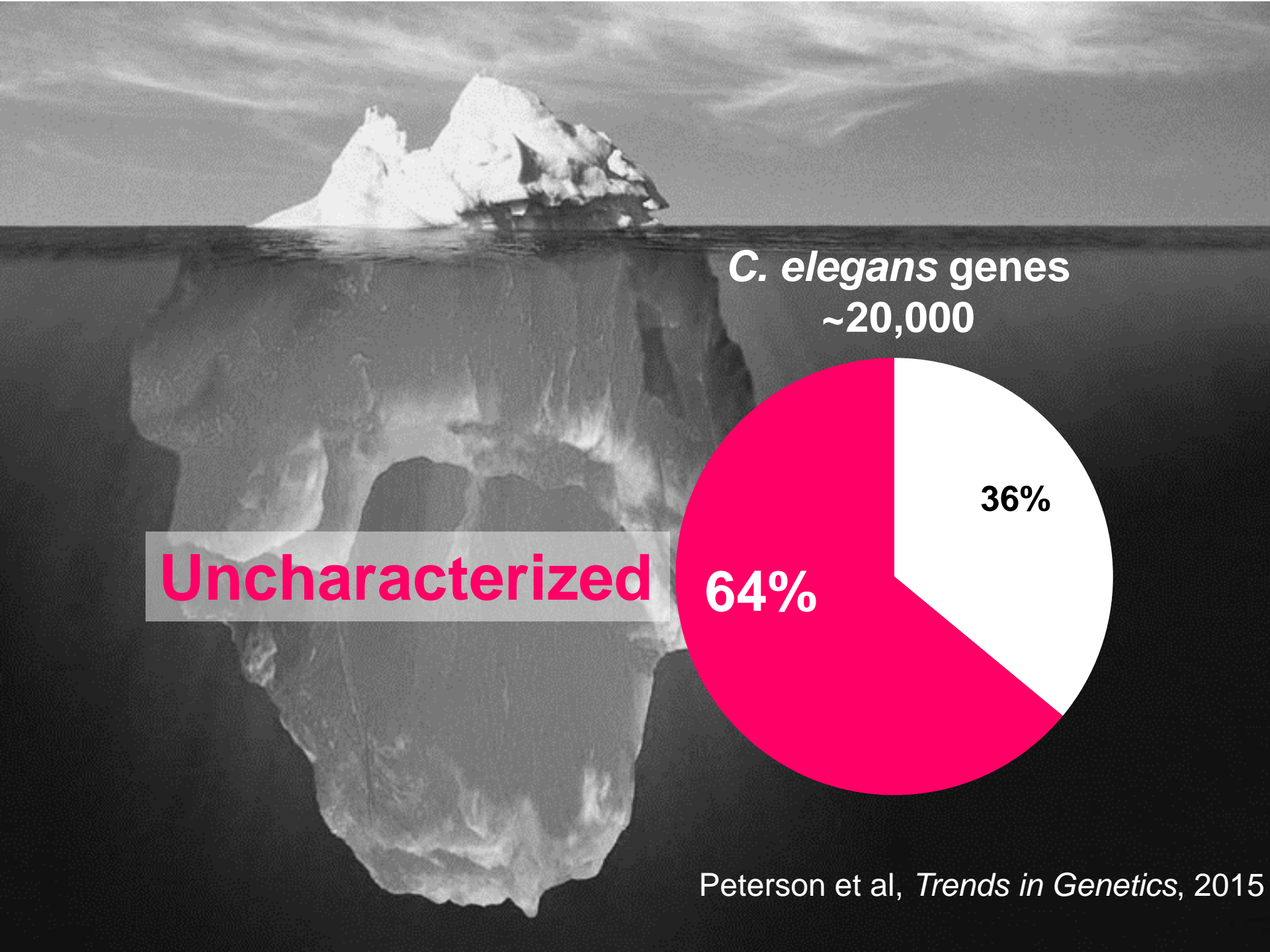
**Worm**



- *Dessert* -

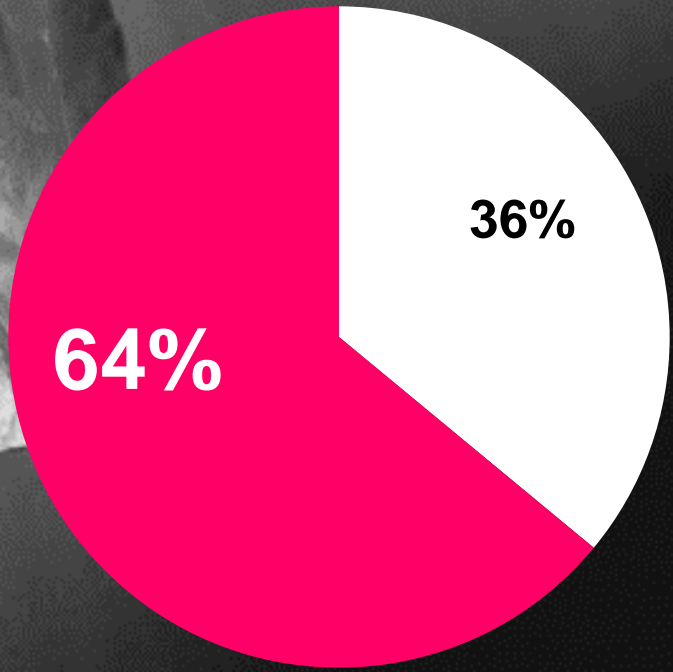
**Me**





*C. elegans* genes  
~20,000

**Uncharacterized**



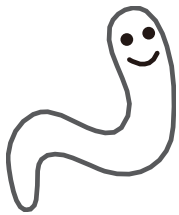
**I want to find and  
characterize new  
genes!!**



**Forward genetic screen**

$$0 + 0 = 1$$

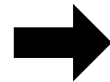
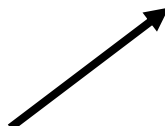
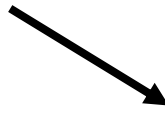
**Blueprint**  
設計図



Gene A

Gene B

Gene C



Gene A

+

Gene B



**Well characterized**

# Combination is challenging

Combination of two genes

$$20,000C_2 = 199,990,000$$



**NEW**  
**Forward genetic screen**

# I can do it with *C. elegans*!



8 elegant things

1. Maintenance

2. Sensation

3. Memory

4. Brain

5. Stereotype

6. Transparency

7. Manipulation

8. Conservation

I hope

you see elegance

in the *C. elegans* brain





# Thanks!

Contact

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2012.10.31 @San Diego, USA