

ADULT CANCER PROGRAM

## Who am I?

### What do I do?



# Clinical Geneticist Cancer Research (colon cancer)

## Who am I?

### What do I do?



Published over 40 research papers describing the genetics and biology of cancer









## Outline





# How can we use a popular concept like zombies to engage and educate

OUTLINE

### What are zombies?



### Impervious to pain

### Aggressive

# Most of these "symptoms" can be explained by brain dysfunction

Instinctive

Undead

Stupid

Can only be killed by destroying the brain

Truncal Ataxia

## What are zombies?



CHARACTERISTICS OF A ZOMBIE?

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Ackermann, H. W.; Gauthier, J. (1991). "The Ways and Nature of the Zombi". <u>The</u> <u>Journal of American Folklore **104** (414):</u> <u>466–494.</u>







## Truncal Ataxia

Stumbling and uncoordinated gait Coordination is controlled by the cerebellum





### Insatiable hunger

Zombies always have the munchies

The region of the brain that controls hunger (satiety) is called the hypothalamus





Specifically, the Ventral Medial Hypothalamus is not receiving signals from gut neurons that they're full

## Stupid!

### Zombies act on instinct

### No cognitive function



Cognition is a function of the frontal lobe



## Aggressive

### Zombies always seem angry

Aggression is controlled by a region of the brain called the amygdala



The amygdala is found deep within the brain in the limbic apparatus





# In normal brains, the frontal lobe and the amygdala communicate





This is where the conscious decision to fight or run comes from

### ZOMBIES DON'T HAVE THIS!

THE SCIENCE BEHIND ZOMBIE CHARACTERISTICS

### Instinctive



# Zombies response to loud noises and movement

Their actions are reflexive



Without the brain stem zombies cease to be







THE SCIENCE BEHIND ZOMBIE CHARACTERISTICS

## Zombiism, or

## Ataxic Neurodegenerative Satiety Deficiency Syndrome (ANSD)

Correct medical name for zombiism!

Ataxic failure of muscular coordination and movement

**Neurodegenerative** degeneration of neurons in the brain

**Satiety** the control of hunger

**Syndrome** A group of symptoms that consistently occur together



Impervious to pain

### "Congenital insensitivity to pain"

Caused by mutations to genes involved in nerve function

Sufferers can bite off parts of their tongue and lips while eating

Rare inherited conditions in which

individuals are unable to perceive pain

Vittangi, Sweden, 40/784 inhabitants

Some individuals were known to perform in "street theatres" by placing knives through arms and walking on burning coals







## Aggression

"The warrior gene"

Aggressive behaviour

Mutations to the MAO-A (monoamine oxidase A) gene involved in nerve function Violence

Impulsivity

Only affects males



## Ataxia

Alzheimer's Disease Dementia Variant Creutzfeldt-Jakob Disease (vCJD) Parkinson's Disease Cerebral Palsy Multiple sclerosis Many other rare diseases Many of these disease damage the brain stem and cerebellum resulting in loss of muscle coordination

Affect speech and movement



### Precedents in Human Disease chickens

# Meet "Mike the headless chicken"

### (a.k.a. Miracle Mike)

Fruita, Colorado, 1945-1947

Mike survived for 18 months after having his head chopped off!

The incomplete decapitation left part of the brain stem

Mike could balance on a perch, walk and attempted to crow and preen his feathers

This is because the brain stem controls breathing, heart rate and reflex actions







#### **BRAIN STEM FUNCTION**



## Sleeping sickness

Prevalent in Africa Caused by the <u>protozoan</u> parasite *Trypanosoma brucei* 

Transmitted by the tsetse fly.



Early-stage symptoms include: Headaches Aching muscles Itching

Late-stage symptoms include:

Irritable, difficulty concentrating, slurred speech, lack of appetite, insomnia at night, narcolepsy during the day,

Coma, death (50k-70k per year)



## **Rabies**

# Transmissible through bites from infected animals such as bats



### Symptoms:

Full or partial paralysis, mental impairment, agitation and strange behaviour, mania, delirium.

55k deaths annually

Incurable once symptoms are present and 99.99 % fatal

Can take several weeks for symptoms while the virus 'migrates' from peripheral neurons to the CNS

Spoiler:

Although rabies causes aggression and attacks in animals, it doesn't in humans



## Necrosis

A condition symptomatic of many different disease including:

Cancer Poisoning Injury

Infection



Cured by "debridement" or amputation

### Disease outbreaks

## Ebola virus

### First described in 1976

### Zaire

318 cases, 280 deaths, 88 % fatality rate

### 28 outbreaks to date

> 1640 deaths, mainly in Africa

Ongoing outbreak in Guinea, Sierra Leone and Guinea

I5I cases, 99 deaths, 66 % fatality rate



### Influenza virus

### Statistics from the US

- 5-20% of population get flu each year
- 200,000 hospitalisations/year
- 3,000 to 49,000 deaths each year from flu-related cause
- Influenza and pneumonia = eighth leading cause of death in males in 2009.

#### New Influenza A (H1N1), Number of laboratory confirmed cases as reported to WHO

### *Status as of 26 June 2009 06:00 GMT*



The boundaries and names shown and the designations used on this map do not imply the expression of any opinion whatsoever on the part of the World Health Organization concerning the legal status of any country, territory, city or area or of its authorities, or concerning the delimitation of its frontiers or boundaries. Dotted lines on maps represent approximate border lines for which there may not yet be full agreement. Data Source: World Health Organization Map Production: Public Health Information and Geographic Information Systems (GIS) World Health Organization



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Map produced: 26 June 2009 07:00 GMT

#### Students learn to:

- 1. What is a healthy organism?
- discuss the difficulties of defining the terms 'health' and 'disease'
- outline how the function of genes, mitosis, cell differentiation and specialisation assist in the maintenance of health

#### Students:

 use available evidence to analyse the links between gene expression and maintenance and repair of body tissues



These differences between cells are controlled by epigenetics



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- Exactly the same genetic information yet....
- Epigenetics determines how and when genes are switched on or off



### Epigenetics controls the way cells read our DNA

THEREEXISTSEVERALDEFINITIONSOFEPIGENETICS ANDASARESULTTHEREAREDISAGREEMENTSASTOWHAT EPIGENETICSSHOULDMEANEPIGENETICCHANGESCAN MODIFYTHEACTIVATIONOFCERTAINGENESBUTNOTTH ESEQUENCEOFDNATHEREFORETHETERMEPIGENETICS REFERSTOFUNCTIONALLYRELEVANTMODIFICATIONS TOTHEGENOMETHATDONOTINVOLVEACHANGEINTHENU CLEOTIDESEQUENCEEXAMPLESOFSUCHMODIFICATIO NSAREDNAMETHYLATIONANDHISTONEMODIFICATION BOTHOFWHICHSERVETOREGULATEGENEEXPRESSIONW ITHOUTALTERINGTHEUNDERLYINGDNASEQUENCESOM ATICEPIGENETICINHERITANCETHROUGHEPIGENETI CMODIFICATIONSPARTICULARLYTHROUGHDNAMETHY LATIONANDCHROMATINREMODELINGISVERYIMPORTA NTINTHEDEVELOPMENTOFMULTICELLULAREUKARYOT ICORGANISMSTHEGENOMESEQUENCEISSTATICWITHS OMENOTABLEEXCEPTIONSBUTCELLSDIFFERENTIATE INTOMANYDIFFERENTTYPESWHICHPERFORMDIFFERE NTFUNCTIONSANDRESPONDDIFFERENTLYTOTHEENVI RONMENTPUNCTUATIONISHUGELYIMPORTANTANDCHA NGESTHEMEANINGOFEVERYTHING

There exist several definitions of epigenetics, and as a result, there are disagreements as to what epigenetics should mean. Epigenetic changes can modify the activation of certain genes, but not the sequence of DNA. Therefore, the term epigenetics refers to functionally relevant modifications to the genome that do not involve a change in the nucleotide sequence. Examples of such modifications are DNA methylation and histone modification, both of which serve to regulate gene expression without altering the underlying DNA sequence. Somatic epigenetic inheritance through epigenetic modifications, particularly through DNA methylation and chromatin remodeling, is very important in the development of multicellular eukaryotic organisms. The genome sequence is static, with some notable exceptions, but cells differentiate into many different types, which perform different functions, and respond differently to the environment.



### Epigenetics controls the way cells **read** our DNA





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### Punctuation is hugely important and can change the meaning of everything!





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The hardware of a computer is useless without the instructions on how to use it

## So how does the cell punctuate our DNA?





Olins and Olins, 2003 Nat Rev Mol Cell Biol. **4**:809



Lieberman-Aiden *et al.* 2009 *Science* **326**:289-293

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These differences between cells are controlled by epigenetics

- 2. Over 3000 years ago the Chinese and Hebrews were advocating cleanliness in food, water and personal hygiene
- distinguish between infectious and non-infectious disease
- explain why cleanliness in food, water and personal <u>hygiene</u> practices assist in control of disease
- identify the conditions under which an organism is described as a pathogen
- identify data sources, plan and choose equipment or resources to perform a first-hand investigation to identify microbes in food or in water
- gather, process and analyse information from secondary sources to describe ways in which drinking water can be treated and use available evidence to explain how these methods reduce the risk of infection from pathogens

### **Definitions**

### Infectious

- An infectious disease is something that can be transmitted.
- Not just humans.
- Usually caused by organisms such as bacteria and viruses but not always

### Pathogen

Something that can cause disease



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### Pathogen?

Not all bacteria and viruses are pathogens

In each of us, our microbial census exceeds the total number of our own human cells by about 10-fold

Symbiotic relationship (e.g. they enhance the energy harvest from our diet)



#### Students learn to:

3. During the second half of the nineteenth century, the work of Pasteur and Koch and other scientists stimulated the search for microbes as causes of disease

- describe the contribution of Pasteur and Koch to our understanding of infectious diseases
- distinguish between:
  - prions
  - viruses
  - bacteria
  - protozoans
    - fungi
  - macro-parasites
  - and name one example of a disease caused by each type of pathogen
- identify the role of antibiotics in the management of infectious disease

#### Students:

- perform an investigation to model Pasteur's experiment to identify the role of microbes in decay
- gather and process information to trace the historical development of our understanding of the cause and prevention of malaria
- identify data sources, gather process and analyse information from secondary sources to describe one named infectious disease in terms of its:
  - cause
  - transmission
  - host response
  - major symptoms
  - treatment
  - prevention
  - control
- process information from secondary sources to discuss problems relating to antibiotic resistance

#### Distinguishing between different infectious agents

#### Bacteria

- Single cell organisms, motile, prokaryotic
- Cell wall
- Antibiotics effective

#### Viruses

- Designed to transmit a nucleic acid genome between h
- Composed of nucleic acid encapsulated in a protein 'capsid'
- Antibiotics ineffective
- Influenza, ebola, rabies, polio,

#### Protozoans

- Single cell organisms, motile, eukaryotic
- Cause parasitic diseases including malaria (plasmodium), intestina and African sleeping sickness (Trypanosoma brucei)

- Proteinacious infection particle
- Infectious agent composed of protein in a misfolded form
- Transmissible spongiform encephalopathies



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### Protozoa





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• Prions cause the molecular equivalent of a zombie apocalypse