

DNAi DVD CONTENTS

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THEMES

• DNA MOLECULE •

Discovery of DNA

A pre-1953 notion	_ biology prior to discovery of the double helix ... François Jacob
DNA is the genetic material	_ the experiment that identified DNA as the genetic material ... Maclyn McCarty
Chargaff's ratios	_ the DNA base ratio rules ... Erwin Chargaff
Race to discover the structure	_ the race between King's College, London, and other groups to define the structure of DNA ... Raymond Gosling
Clue: position of phosphates	_ realizing phosphates are on the outside of the structure ... Raymond Gosling
An earlier DNA model	_ Franklin's analysis of Watson and Crick's early model of DNA ... Raymond Gosling
Clue: X-ray diffraction	_ how the X-ray diffraction camera works ... Raymond Gosling
	_ the X-ray diffraction picture that revealed the helix ... Maurice Wilkins
The answer	_ working out the structure of DNA ... Jim Watson
DNA: the secret of life	_ on his and Francis Crick's gigantic breakthrough ... Jim Watson
DNA: the key to understanding	_ why the discovery of DNA's structure was so important ... Francis Crick

Structure of DNA

An elegant structure	_ the elegant simplicity of Watson and Crick's model ... Raymond Gosling
Replication models	_ the different models proposed for DNA replication ... Matthew Meselson
The correct model	_ Meselson and Franklin Stahl's experiment to determine the correct DNA replication mode ... Matthew Meselson
Replicating the helix	_ animation
Mechanism of replication	_ animation
How DNA is packaged	_ animation

Organization of DNA

Billions of bases	_ there are 2.9 billion letters in the human genome ... Mark Adams
Chromosome map	_ animation
100 km of DNA	_ the length of the human genome ... John Sulston
Walking down a chromosome	_ traveling down the genome ... Ewan Birney
Coding vs. non-coding	_ how much of the genome is active? ... Jim Kent
How much DNA codes for protein?	_ animation
An important gene cluster	_ a cluster of immunity genes on chromosome six ... Jim Kent
Bacterial vs. human genome	_ human genes are organized in patches of information ... Eric Lander

• DNA IN ACTION •

The genetic code

- The digital code
 - The coding problem
 - DNA has four units
 - Defining the gene
 - 3 DNA bases = 1 amino acid
 - Code analogies
 - Triplet code
 - Cracking the first codon
 - Cracking the code
 - Universal code
- _ DNA is a digital code that can be read ... Leroy Hood
 - _ the problem posed by Watson and Crick's model ... Sydney Brenner
 - _ animation
 - _ matching the gene to protein sequence ... Sydney Brenner
 - _ the mathematician George Gamow's idea that three DNA bases encode one amino acid ... Marshall Nirenberg
 - _ the idea of applying a code to DNA sequence ... Sydney Brenner
 - _ animation
 - _ deciphering the first amino acid codon ... Marshall Nirenberg
 - _ deciphering every triplet code ... Marshall Nirenberg
 - _ all forms of life use the same genetic instructions ... Marshall Nirenberg

The Central Dogma & gene expression

- The Central Dogma & gene expression
 - Protein synthesis
 - The role of the ribosome
 - Transcription
 - Explaining the Central Dogma
 - RNA splicing
 - Processing mRNA
 - Translation
 - Gene regulation
 - The lac operon model
- _ DNA makes RNA makes protein ... Jim Watson
 - _ synthetic RNA stimulates protein synthesis ... Marshall Nirenberg
 - _ ribosomes recognize a triplet code ... Marshall Nirenberg
 - _ animation
 - _ the mechanism of protein synthesis and the virus (phage) experiment that proved it ... Sydney Brenner
 - _ RNA is spliced ... Tom Cech
 - _ mRNA editing by the spliceosome ... Eric Lander
 - _ animation
 - _ the regulation of genes by control proteins ... Walter Gilbert
 - _ his model for bacterial gene regulation ... François Jacob

Mutation & evolution

- Random mutations
 - Sickle cell
 - DNA damage
 - Junk DNA & evolution
 - Humans & chimps
 - Regulatory networks
 - Differences & similarities
 - Evolutionary relationships
 - Smell receptors
 - What came first: DNA or RNA?
- _ random mutations are necessary for evolution ... Jim Kent
 - _ animation
 - _ animation
 - _ junk DNA may have important evolutionary functions ... Eric Lander
 - _ humans and chimps share around 99% of their DNA ... Mary-Claire King
 - _ the importance of regulatory systems for evolution ... Leroy Hood
 - _ DNA variations result in differences between individuals ... Robert Plomin
 - _ the conservation of life processes ... John Sulston
 - _ every gene has a distinctive evolutionary history ... Eric Lander
 - _ evidence that RNA evolved before DNA ... Tom Cech

• GENETICS TO GENOMICS •

Key experiments & techniques

- Polymerase chain reaction
 - DNA variation
 - DNA fingerprinting
 - First recombinant DNA
 - Mechanism of recombination
 - Microarray analysis
 - DNA transfer: agrobacterium
 - DNA transfer: gene gun
- _ discovery of the PCR technique ... Kary Mullis
 - _ measuring DNA variation: techniques and applications ... Mark Skolnick
 - _ using minisatellites (tandem DNA repeats) to create unique genetic profiles ... Alec Jeffreys
 - _ describing the first experiment with recombinant DNA ... Paul Berg
 - _ animation
 - _ studying gene expression using microarrays ... Pat Brown
 - _ transferring genes into plant cells using agrobacterium ... Robert Horsch
 - _ inserting genes into plant cells using a gene gun ... Robert Horsch

Key experiments & techniques

Sequencing DNA
Sequencing genomes

- (... continued)
_ inside a DNA sequencing machine ... Leroy Hood
_ the speed of sequencing since automation ... Mike Hunkapiller

Advances & applications

Cross-species recombination
Birth of genetic engineering

The impact of cloning
Cloning DNA in bacteria
Risks of DNA recombination
Lab safety

- _ first experiment to recombine DNA from different species ... Stanley Cohen
_ significance of his experiment with Stanley Cohen to clone toad DNA ... Herbert Boyer
_ the implications of cloning mammalian genes ... Herbert Boyer
_ importance of being able to clone DNA using bacteria ... Paul Berg
_ potential risks associated with recombining DNA ... Robert Pollack
_ demonstrating the P4 lab containment suit he developed for working with high risk substances ... Emmett Barkley
_ the Genentech method of producing ... David V. Goeddel
_ the first transgenic crop, engineered by Monsanto ... Robert Horsch
_ cotton plants engineered to be pest resistant ... Jim Watson
_ raising concerns associated with GM crop production ... Jim Kent

Insulin production
First transgenic crop
Cotton plants
GM crop concerns

The Human Genome Project

The aim
The motivation
Public & private
Public project sequencing
Assembling the fragments
Private project sequencing
Reading the genome
The completed genome
Outcome of the HGP
After the HGP

- _ the aim of the Human Genome Project ... Jim Watson
_ the justification for the Human Genome Project ... Francis Collins
_ comparing methods used by the public and private teams ... Gene Myers
_ animation
_ problems assembling the genome fragments ... Jim Kent
_ animation
_ interpreting the completed human genome sequence ... Ewan Birney
_ the completion of the draft human genome sequence ... William J. Clinton
_ a new paradigm for studying biology ... Eric Lander
_ a new foundation for science ... J. Craig Venter

Bioinformatics

An overview
Solving a problem
Computing power
Analyzing your genes
Proteomics
Implications for the future

- _ using computers to assemble genomes and interpret data ... Gene Myers
_ developing the tools to sequence the genome ... J. Craig Venter
_ computational power of a processing farm ... Ewan Birney
_ understanding the genome will lead to medical advances ... Leroy Hood
_ studying proteins to understand disease ... Scott Patterson
_ new tools for redesigning life ... Leroy Hood

• HUMAN GENETICS •

Disease research

The challenge of gene hunting
Locating disease genes
Impact of the genome projects
Animal models
The DMD approach

- _ the challenge of finding a disease gene ... Francis Collins
_ locating disease genes using markers ... David Botstein
_ the increased speed of gene searching ... Ewan Birney
_ using mouse models to study disease ... Mario Capecchi
_ gene replacement therapy in Duchenne muscular dystrophy (DMD) ... Kay Davies
_ gene expression patterns in diseased cells ... Pat Brown
_ switching genes on and off to study disease ... Mario Capecchi
_ using embryonic stem cells to make mouse models ... Mario Capecchi

Gene expression patterns
Gene switches
Gene manipulation

Disease research

Living with sickle cell
Inheriting sickle cell

(... continued)

_ how sickle cell has affected her life ... Katreece McGhee
_ how she inherited sickle cell ... Katreece McGhee

Work on cancer

Mutations & cancer
Tumor growth
Cancer genes
Early cancer studies
Finding cancer genes

_ cancer is caused by an accumulation of mutations ... Bruce Ames
_ animation
_ describing tumor suppressors and oncogenes ... Mike Wigler
_ studying cancer prior to understanding its mechanisms ... Mary-Claire King
_ searching for candidate genes in families with breast cancer
... Mary-Claire King
_ identifying and tracking genetic markers using family trees ... Barbara Weber
_ finding and cloning the first breast cancer gene: BRCA1 ... Mark Skolnick
_ finding the second breast cancer gene: BRCA2 ... Mark Skolnick
_ current status of testing for cancer genes ... Mary-Claire King
_ hopes for cancer treatments ... Mary-Claire King

Using family trees
Identifying BRCA1
Looking for BRCA2
Limitations of testing
Hopes for the future

Screening & treatment

Leukemia: the Gleevec story
How Gleevec works
Gleevec: first trials
Developing other cancer drugs
More questions than answers
Genetic screening
Offering options
Testing for a reason

_ the development of Gleevec, a drug to treat leukemia ... Brian J. Druker
_ animation
_ the first patient in the Gleevec trials ... Bud and Yvonne Romine
_ applying the Gleevec model to other cancers ... Brian J. Druker
_ approaching population screening with caution ... Francis Collins
_ setting up a screen for muscular dystrophy ... Kay Davies
_ importance of choice regarding genetic testing ... Kay Davies
_ schizophrenia: a case for testing ... Kay Jamison

Behavioral genetics

Genetic pre-wiring
Nature vs. nurture
Complex behavior
Twin studies
Prediction & prevention
Heritability of behaviors
Mental illness & creativity
Basis of complex disorders

_ behavior can be both genetically pre-wired and learnt ... Hubert Markl
_ how much of our behavior can be attributed to genes? ... Robert Plomin
_ the honeybee as a model for complex behavior ... Hubert Markl
_ the genetic basis of cognitive traits ... Robert Plomin
_ predicting and preventing behavioral problems ... Robert Plomin
_ all behavioral traits have a heritable component ... Robert Plomin
_ genetic links between mental illness and creativity ... Kay Jamison
_ understanding the genetic basis of complex traits ... Robert Plomin

Human origins

Neandertal DNA

Neandertal & human ancestry
Counting DNA mutations

Tracking human history

The evolutionary puzzle
The divergence of Neandertals

A recent common ancestor

Classification and value

_ comparing Neandertal and modern human mitochondrial DNA
... Svante Pääbo
_ human origins and our common ancestry with Neandertals ... Svante Pääbo
_ why the number of mutations in mitochondrial DNA is an underestimate
... Mark Stoneking
_ using the Y chromosome and other genomic regions to track human history
... Michael F. Hammer
_ genetic data must be part of a framework ... Michael F. Hammer
_ fossil evidence shows that Neandertals diverged from modern humans
... Chris Stringer
_ mitochondrial DNA confirms a recent common ancestor for modern humans
... Douglas Wallace
_ confounding genetic classification with human worth ... Hubert Markl

• ETHICS & IMPLICATIONS •

Eugenics

Sterilization as a welfare reform
Pre-WWII German eugenics
American perspective
The Buck vs. Bell case
Reactions to imperfections
Who should decide?
Directing our evolution
Classification and value

_ sterilization in the USA: used as a welfare reform ... Paul Lombardo
_ eugenics in Nazi Germany ... Jim Watson
_ an overview of eugenics in the USA ... Jim Watson
_ sterilization in the USA: Buck vs. Bell ... Paul Lombardo
_ should we correct natural genetic imperfections? ... Jim Watson
_ making life choices and economic considerations ... Benno Müller-Hill
_ our responsibility to direct our own evolution ... Jim Watson
_ confounding genetic classification with human worth ... Hubert Markl

Diversity & enhancement

None of us are perfect
Vanilla children
Designer babies
Weeding out disease
Harrington family
A case for testing
Protecting diversity
A better understanding

_ human imperfections and genetic enhancement ... Jim Watson
_ for diversity and against narrowing the options ... Kay Jamison
_ intervening in a child's future at a genetic or social level ... Robert Plomin
_ predictions for gene testing ... Bruce Ames
_ his relationship with his son who has Down syndrome ... Roby Harrington
_ testing and managing genetic disorders ... Kay Davies
_ manic depressives: an endangered but valued species ... Kay Jamison
_ improving our species with better education ... Hubert Markl

Ownership & access

Patenting living organisms
Selecting genes to patent
Patenting chaos
Commercial patents
Human genome patents
A free flow of information

_ creating and owning living organisms ... Ananda Chakrabarty
_ a private company's approach to patenting genes ... Mark Adams
_ on knowing the function of a gene before you patent ... Mary-Claire King
_ on patenting genes for commercial purposes ... Mary-Claire King
_ the human genome sequence is not a basis for a patent ... John Sulston
_ making sequence public to pre-empt the patents ... John Sulston

• REFLECTING ON SCIENCE •

A selection of views

Science & faith
Playing God
Explaining life through science
Reading our own code
Influencing our evolution
Manic depression
Definitions of life
Can genetics provide answers?

_ reconciling working in science with faith in God ... Francis Collins
_ miracles from knowledge, not prayer ... Jim Watson
_ reading a letter from Rosalind to her father ... Rosalind Franklin's sister
_ reflecting on our evolution ... John Sulston
_ eliminating faults in our genetic programming ... Raymond Gosling
_ mental illness and complicated choices ... Kay Jamison
_ defining what it is to be alive ... Tom Cech
_ genetics may not provide the answers we seek ... Benno Müller-Hill

Still to be explored

Unused tools
Future of medicine
Manipulating living systems
Germline therapy
Redesigning organisms
The future of humans

_ gene technology and its possible uses ... Robert Pollack
_ predictive and preventative personalized medicine ... Leroy Hood
_ manufacturing new proteins in living systems ... Robert Horsch
_ on needing to make germline therapy reversible ... Mario Capecchi
_ technological advances may allow us to redesign life ... Leroy Hood
_ what will humans look like in 5,000 years? ... Ananda Chakrabarty

ANIMATIONS

DNA molecule

- DNA has four units
- Chargaff's ratios
- Triple helix
- Base pairing
- How DNA is packaged *
- DNA unzip
- Chromosome map
- How much DNA codes for protein?

Replication

- Replicating the helix
- Mechanism of replication *

Transcription & translation

- Triplet code
- Transcription *
- mRNA splicing
- Translation *

Experiments & techniques

- Mechanism of recombination
- Microarray
- Polymerase chain reaction
- Sanger sequencing
- Public project sequencing
- Private project sequencing

Disease & mutation

- DNA damage
- Sickle cell
- Tumor growth
- How Gleevec works

(*) narration options: none, basic, and advanced.

INTERVIEWS

1)

Mark Adams

- Billions of bases
- Selecting genes to patent

Bruce Ames

- Mutations & cancer
- Weeding out disease

Emmett Barkley

- Lab safety

Paul Berg

- First recombinant DNA
- Cloning DNA in bacteria
- Cohesive ends & recombination

Ewan Birney

- Walking down a chromosome
- Reading the genome
- Computing power
- Impact of the genome projects

David Botstein

- Locating disease genes
- Cost of the Human Genome Project
- Opposition to the Human Genome Project

Herbert Boyer

- Birth of genetic engineering
- The impact of cloning
- Why study plasmids?

Sydney Brenner **

- The coding problem
- Defining the gene
- Code analogies
- Explaining the Central Dogma
- The coiled nature of DNA
- Cell organization

Pat Brown

- Microarray analysis
- Gene expression patterns

Mario Capecchi

- Animal models
- Gene switches
- Gene manipulation
- Germline therapy

2)

Tom Cech **

RNA splicing
What came first: DNA or RNA?
Definitions of life

Ananda Chakrabarty

Patenting living organisms
The future of humans

Erwin Chargaff

Chargaff's ratios

William J. Clinton

The completed genome

Stanley Cohen

Cross-species recombination

Francis Collins

The motivation (HGP)
The challenge of gene hunting
More questions than answers
Science & faith

Francis Crick **

DNA: the key to understanding
The fascination of science
DNA's deceptive "simplicity"
Understanding the brain

Kay Davies

The DMD approach
Genetic screening
Offering options
A case for testing

Brian J. Druker

Leukemia: the Gleevec story
Developing other cancer drugs

Rosalind Franklin's sister

Explaining life through science

3)

Walter Gilbert **

Gene regulation
Experiments to find RNA
The repressor/inducer system

David V. Goeddel

Insulin production

Raymond Gosling

Race to discover the structure
Clue: position of phosphates
An earlier DNA mode
Clue: X-ray diffraction
An elegant structure
Influencing our evolution

Michael F. Hammer

Tracking human history
The evolutionary puzzle

Roby Harrington

Harrington family

Leroy Hood

The digital code
Regulatory networks
Sequencing DNA
Analyzing your genes
Implications for the future
Future of medicine
Redesigning organisms

Robert Horsch

DNA transfer: agrobacterium
DNA transfer: gene gun
First transgenic crop
Manipulating living systems

Mike Hunkapiller

Sequencing genomes

François Jacob **

A pre-1953 notion
The lac operon model

Kay Jamison

Testing for a reason
Mental illness & creativity
Vanilla children
Protecting diversity
Manic depression

4)

Alec Jeffreys

DNA fingerprinting

Jim Kent

Coding vs. non-coding
An important gene cluster
Random mutations
GM crop concerns
Assembling the fragments

Mary-Claire King

- Humans & chimps
- Early cancer studies
- Finding cancer genes
- Limitations of testing
- Hopes for the future
- Patenting chaos
- Commercial patents

Arthur Kornberg **

- Enzymes: DNA polymerase
- Studying DNA replication
- DNA synthesis

Eric Lander

- Bacterial vs. human genome
- Processing mRNA
- Junk DNA & evolution
- Smell receptors
- Outcome of the HGP

Paul Lombardo

- Sterilization as a welfare reform
- The Buck vs. Bell case

Hubert Markl

- Genetic pre-wiring
- Complex behavior
- Classification and value
- A better understanding

Maclyn McCarty

- DNA is the genetic material

Katreece McGhee

- Living with sickle cell
- Inheriting sickle cell

Matthew Meselson

- Replication models
- The correct model

5)

Benno Müller-Hill

- Who should decide?
- Can genetics provide answers?

Kary Mullis **

- Polymerase chain reaction

Gene Myers

- Public & private
- An overview (bioinformatics)

Marshall Nirenberg **

- 3 DNA bases = 1 amino acid
- Cracking the first codon
- Cracking the code
- Universal code
- Protein synthesis
- The role of the ribosome

Svante Pääbo

- Neandertal DNA
- Neandertal & human ancestry

Scott Patterson

- Proteomics

Robert Plomin

- Differences & similarities
- Nature vs. nurture
- Twin studies
- Prediction & prevention
- Heritability of behaviors
- Basis of complex disorders
- Designer babies

Robert Pollack

- Risks of DNA recombination
- Unused tools

Bud and Yvonne Romine

- Gleevec: first trials

Mark Skolnick

- DNA variation
- Identifying BRCA1
- Looking for BRCA2

6)

Mark Stoneking

- Counting DNA mutations

Chris Stringer

- The divergence of Neandertals

John Sulston **

- 100 km of DNA
- Evolutionary relationships
- Human genome patents
- A free flow of information
- Reading our own code

J. Craig Venter

- After the Human Genome Project
- Solving a problem

Douglas Wallace

A recent common ancestor

Jim Watson **

The answer
DNA: the secret of life
Cotton plants
The aim (Human Genome Project)
Need for an RNA template
Pre-WWII German eugenics
Reactions to imperfections
Directing our evolution
None of us are perfect
American perspective
Playing God

Barbara Weber

Using family trees

Mike Wigler

Cancer genes

Maurice Wilkins **

Clue: X-ray diffraction

(**) Nobel Laureate

BACKGROUND BRIEFING

Animator: Drew Berry

Making 3D animations
Working with scientists
Using 3D models
Absolute accuracy
A complex animation

_ the process of making 3D scientific animations
_ science as a creative career
_ online libraries of molecules can model interactions
_ ensuring the scientific accuracy of the models
_ creating the replication animation

Teacher: Caren Gough

Teacher involvement
Using the DVD
Planning lessons
Interviews on the DVD
Ease of use

_ teacher contribution to content selection
_ planning lessons using the DVD content
_ planning lessons using the animations
_ scientist interviews provide a unique resource
_ on expertise required to use the DVD