DNAi DVD and the DNAi Teacher Guide

The *DNA Interactive (DNAi)* DVD carries approximately four hours of video interviews with 11 Nobel Laureates and more than 50 other scientists, clinicians, and patients. It also holds the complete set of 3-dimensional animations produced for the *DNA* TV series and *DNAi* project.

The following pages list video clips and animations from the DVD that would be appropriate to show with specific activities in the *DNAi Teacher Guide*. The clips and animations are listed under "themes" and "additional animations." The "themes" listing includes relevant interviews and animations that can be accessed from the "themes" section of the DVD. The "additional animations" are best accessed from "animations" button in the DVD main menu.

You can access the DNAi Teacher Guide by registering at www.dnai.org/teacher.

Activity 1: DNAi Timeline: a scavenger hunt

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
The answer	_ the X-ray diffraction picture that revealed the helix Maurice Wilkins
DNA: the key to understanding	_ why the discovery of DNA's structure was so important Francis Crick
Structure of DNA	
The correct model	_ Meselson and Franklin Stahl's experiment to determine the correct DNA replication mode Matthew Meselson
• DNA IN ACTION •	
The genetic code	
Defining the gene	_ matching the gene to protein sequence Sydney Brenner
Cracking the first codon	_ deciphering the first amino acid codon Marshall Nirenberg
The Central Dogma and gene expre	
Explaining the Central Dogma	_ the mechanism of protein synthesis and the virus (phage) experiment that proved it Sydney Brenner
The lac operon model	_ his model for bacterial gene regulation François Jacob
• GENETICS TO GENOMICS •	
Key experiments and techniques	
Polymerase chain reaction	_ discovery of the PCR technique Kary Mullis
DNA fingerprinting	_ using minisatellites (tandem DNA repeats) to create unique genetic profiles Alec Jeffreys
First recombinant DNA	_ describing the first experiment with recombinant DNA Paul Berg
Microarray analysis	_ studying gene expression using microarrays Pat Brown

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Advances and applications

Cross-species recombination

Birth of genetic engineering

<u>The Human Genome Project</u> The completed genome

• HUMAN GENETICS •

Disease research Gene manipulation

<u>Work on cancer</u> Finding cancer genes

- _ first experiment to recombine DNA from different species . . . Stanley Cohen
- _ significance of his experiment with Stanley Cohen to clone toad DNA ... Herbert Boyer

_ the completion of the draft human genome sequence . . . William J. Clinton

_ using embryonic stem cells to make mouse models . . . Mario Capecchi

_ searching for candidate genes in families with breast cancer ... Mary-Claire King

Activity 2. Finding the Structure: pieces of the puzzle

THEMES

• DNA MOLECULE •

Discovery of DNA

A pre-1953 notion DNA is the genetic material

Chargaff's ratios Race to discover the structure

Clue: position of phosphates

An earlier DNA model

Clue: X-ray diffraction The answer DNA: the secret of life DNA: the key to understanding

ADDITIONAL ANIMATIONS

DNA molecule

DNA has four units Chargaff's ratios Triple helix Base pairing

- $_$ biology prior to discovery of the double helix \ldots François Jacob
- _ the experiment that identified DNA as the genetic material . . . Maclyn McCarty
- _ the DNA base ratio rules ... Erwin Chargaff
- _ the race between King's College, London, and other groups to define the structure of DNA ... Raymond Gosling
- _ realizing phosphates are on the outside of the structure . . . Raymond Gosling
- _ Franklin's analysis of Watson and Crick's early model of DNA ... Raymond Gosling
- _ how the X-ray diffraction camera works ... Raymond Gosling
- _ the X-ray diffraction picture that revealed the helix ... Maurice Wilkins
- _ on his and Francis Crick's gigantic breakthrough ... Jim Watson
- _ why the discovery of DNA's structure was so important . . . Francis Crick



Activity 3. (Clues to) Copying the Code: examining the evidence

_ animation

_ animation

_ the elegant simplicity of Watson and Crick's model . . . Raymond Gosling

the different models proposed for DNA replication ... Matthew Meselson

_ Meselson and Franklin Stahl's experiment to determine the correct DNA

replication mode ... Matthew Meselson

THEMES

• DNA MOLECULE •

Structure of DNA

An elegant structure Replication models The correct model

Replicating the helix Mechanism of replication

ADDITIONAL ANIMATIONS

DNA molecule

DNA unzip **Replication** Replicating the helix Mechanism and replication

Activity 4. Reading the Code

THEMES

• DNA IN ACTION •

The genetic code

The digital code	_ DNA is a digital code that can be read Leroy Hood
The coding problem	_ the problem posed by Watson and Crick's model Sydney Brenner
DNA has four units	_ animation
Defining the gene	_ matching the gene to protein sequence Sydney Brenner
3 DNA bases = 1 amino acid	_ the mathematician George Gamow's idea that three DNA bases encode one
	amino acid Marshall Nirenberg

The Central Dogma & gene expression

Need for an RNA Template	_ DNA makes RNA makes protein Jim Watson
Protein synthesis	_ synthetic RNA stimulates protein synthesis Marshall Nirenberg
The role of the ribosome	_ ribosomes recognize a triplet code Marshall Nirenberg
Transcription	_ animation
Explaining the Central Dogma	_ the mechanism of protein synthesis and the virus (phage) experiment that
	proved it Sydney Brenner
RNA splicing	_ RNA is spliced Tom Cech
Processing mRNA	_ mRNA editing by the spliceosome Eric Lander
Translation	_ animation

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ADDITIONAL ANIMATIONS

DNA molecule

How much DNA codes for protein

Activity 5. Controlling the Code: molecules at work

THEMES

• DNA IN ACTION •

The Central Dogma & gene epxression

Gene regulation The lac operon model _ every gene has a distinctive evolutionary history ... Eric Lander _ evidence that RNA evolved before DNA ... Tom Cech

ADDITIONAL ANIMATIONS

Transcription and translation Triplet code Transcription Translation Transcription and translation How DNA is packaged

Activity 6. Manipulation: transferring genes

THEMES

• 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 Sequencing DNA Sequencing genomes _ 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
- _ 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

Insulin production First transgenic crop Cotton plants GM crop concerns

- _ 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 $\ldots\,$ 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 \hdots . . . 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 \ldots .Jim Kent

ADDITIONAL ANIMATIONS

Experiments and techniques

Polymerase chain reaction Sanger sequencing

Activity 7. Gel Electrophoresis: sort and see the DNA

THEMES

GENETICS TO GENOMICS

<u>Key experiments & techniques</u> DNA fingerprinting

_ using mini satellites (tandem DNA repeats) to create unique genetic profiles . . . Alec Jeffreys

Activity 8. Model Organisms: the genes we share

THEMES

• DNA IN ACTION •

The genetic code

Universal code

Mutation and evolution

Random mutations Junk DNA & evolution Evolutionary relationships $_$ all forms of life use the same genetic instructions . . . Marshall Nirenberg

- _ random mutations are necessary for evolution ... Jim Kent
- __junk DNA may have important evolutionary functions ... Eric Lander
- _ the conservation of life processes ... John Sulston



• GENETICS TO GENOMICS •

Key Experiments & techniques

Sequencing genomes

• HUMAN GENETICS •

Disease research

The challenge of gene hunting Locating disease genes Animal models Gene manipulation

Human origins

The evolutionary puzzle

ADDITIONAL ANIMATIONS

Disease and mutation DNA damage

Activity 9. Sequencing

THEMES

• GENETICS TO GENOMICS •

Key experiments and techniques

Sequencing DNA Sequencing genomes

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

ADDITIONAL ANIMATIONS

Experiments and techniques

Mechanism and recombination Sanger sequencing _ the speed of sequencing since automation \dots Mike Hunkapiller

- _ the challenge of finding a disease gene ... Francis Collins
- _ locating disease genes using markers ... David Botstein
- _ using mouse models to study disease ... Mario Capecchi
- _ using embryonic stem cells to make mouse models ... Mario Capecchi

_ genetic data must be part of a framework ... Michael F. Hammer

- _ inside a DNA sequencing machine ... Leroy Hood
- _ the speed of sequencing since automation ... Mike Hunkapiller
- _ 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 \hdots Jim Kent
- _ animation
- _ interpreting the completed human genome sequence \ldots Ewan Birney
- $_$ the completion of the draft human genome sequence . . . William J. Clinton
- $_$ a new paradigm for studying biology $\ldots\,$ Eric Lander
- _ a new foundation for science ... J. Craig Venter





Activity 10. Genome: a tour and genetic disorder brochure

THEMES

• DNA MOLECULE •

Organization of DNA

Billions of bases Chromosome map 100 km of DNA Walking down a chromosome Coding vs. non-coding How much DNA codes for protein? An important gene cluster Bacterial vs. human genome

• HUMAN GENETICS •

Disease research

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

Gene expression patterns Gene switches Gene manipulation Living with sickle cell Inheriting sickle cell

Work on cancer

Mutations & cancer Tumor growth Cancer genes Early cancer studies

Finding cancer genes

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

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- _ there are 2.9 billion letters in the human genome ... Mark Adams _ animation
- _ the length of the human genome ... John Sulston
- _ traveling down the genome ... Ewan Birney
- how much of the genome is active? ... Jim Kent
- _ animation
- _ a cluster of immunity genes on chromosome six ... Jim Kent
- _ human genes are organized in patches of information . . . Eric Lander
- _ 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
- how sickle cell has affected her life ... Katreece McGhee
- how she inherited sickle cell ... Katreece McGhee
- $_$ cancer is caused by an accumulation of mutations \ldots 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

_ 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 $\ldots\,$ Francis Collins



Genetic screening Offering options Testing for a reason

ADDITIONAL ANIMATIONS

Disease and mutation

DNA damage Sickle cell Tumor growth How Gleevec works **DNA molecule** How DNA is packaged **Experiments and techniques** Microarray Polymerase chain reaction Sanger sequencing

Activity 11. Genome Mining

THEMES

• GENETICS TO GENOMICS •

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

Bioinformatics

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

ADDITIONAL ANIMATIONS

Experiments and techniques

Microarray Sanger sequencing **DNA molecule** DNA unzip How much DNA codes for protein?

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_ setting up a screen for muscular dystrophy ... Kay Davies
 _ importance of choice regarding genetic testing ... Kay Davies
 _ schizophrenia: a case for testing ... Kay Jamison

- _ 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 \hdots . . . 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
- _ 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



THEMES

• DNA IN ACTION •

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?

• HUMAN GENETICS •

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

ADDITIONAL ANIMATIONS

Transcription and translation

Transcription mRNA splicing Translation **Disease and mutation** DNA damage

- $_$ random mutations are necessary for evolution \ldots . 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 $\ldots\,$ Eric Lander
- $_$ evidence that RNA evolved before DNA $\ldots\,$ Tom Cech
- _ 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



Activity 13. Recovering the Romanovs

THEMES

• MUTATION & EVOLUTION •

Random mutations DNA damage Differences & similarities	 _ random mutations are necessary for evolution Jim Kent _ animation _ DNA variations result in differences between individuals Robert Plomin
• HUMAN ORIGINS •	
Counting DNA mutations	_ why the number of mutations in mitochondrial DNA is an underestimate Mark Stoneking

... Michael F. Hammer

Tracking human history

The evolutionary puzzle

ADDITIONAL ANIMATIONS

Experiments and techniques PCR Sanger sequencing

Activity 14. Dealing with DNA controversy: issues, arguments, and ethics

THEMES

• 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

Diversity & enhancement

None of us are perfect Vanilla children Designer babies Weeding out disease Harrington family __sterilization in the USA: used as a welfare reform ... Paul Lombardo eugenics in Nazi Germany ... Jim Watson

using the Y chromosome and other genomic regions to track human history

_ genetic data must be part of a framework ... Michael F. Hammer

- 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
- _ 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



A case for testing Protecting diversity A better understanding

Ownership & access

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

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?

Still to be explored

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

- _ testing and managing genetic disorders ... Kay Davies
- _ manic depressives: an endangered but valued species ... Kay Jamison
- _ improving our species with better education ... Hubert Markl
- _ 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
- _ 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
- _ 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? . . . An and a Chakrabarty

