



The Making of the Fittest: Natural Selection and Adaptation

LESSON
STUDENT HANDOUT

MOLECULAR GENETICS OF COLOR MUTATIONS IN ROCK POCKET MICE

INTRODUCTION

THE ROCK POCKET MOUSE

The rock pocket mouse, *Chaetodipus intermedius*, is a small, nocturnal animal found in the deserts of the southwestern United States. Because most rock pocket mice have a sandy, light-colored coat, they are able to blend in with the light color of the desert rocks and sand that they live on. But populations of primarily dark-colored rock pocket mice have been found living in areas where the ground is covered in a dark rock called basalt, which was caused by geologic lava flows thousands of years ago. Scientists have collected data from a population of primarily dark-colored mice living in an area of basalt in Arizona's Pinacate lava flow, as well as from a nearby light-colored population. Researchers analyzed the data from these two populations to search for the genetic mutation responsible for the dark coat color. Through their analyses, they discovered a mutation in the *Mc1r* gene that is involved in coat-color determination.

THE *MC1R* GENE

The coat color of rock pocket mice is primarily determined by two pigments: **eumelanin**, which is dark colored, and **pheomelanin**, which is light colored. The synthesis of these pigments is controlled by the products of several genes, including the *Mc1r* gene. This gene encodes a protein called melanocortin 1 receptor (MC1R). This receptor is found embedded in the membrane of melanocytes, which are cells specialized for pigment production. The melanocytes of **wild-type** (nonmutant) mice produce more pheomelanin than eumelanin. The result is a **sandy-colored mouse**. The **mutated** version of the *Mc1r* gene, however, triggers melanocytes to increase the production of eumelanin, resulting in the **dark coat-color phenotype**.

GENE MUTATION

A **gene mutation** is any change in the DNA sequence of a gene. Gene mutations can change the structure of the resulting protein. A change in protein structure can change, negate, or have no effect on function. There are several types of mutations, and several results that mutations can have on the amino acid sequences of proteins.

Types of Mutations

- **Substitution mutation:** The replacement of one nucleotide of DNA for another. Mutations that affect a single nucleotide are called point mutations.
- **Insertion mutation:** The addition of one or more nucleotides to the DNA gene sequence. The insertion of nucleotides can result in frameshift mutations.
- **Deletion mutation:** The loss of one or more nucleotides from the DNA gene sequence. The deletion of nucleotides can result in frameshift mutations.

Potential Results a Gene Mutation Has on a Protein

- **Silent mutation:** This mutation does not cause a change in the amino acid sequence of the protein; therefore, there is *no* change in the resulting protein.
- **Missense mutation:** This mutation causes an amino acid in the sequence to be changed to another amino acid. This type of mutation causes a change in the primary structure of the protein (the linear sequence of amino acids), which typically results in a change in the three-dimensional conformation of the protein.
- **Nonsense mutation:** This mutation causes the protein to be truncated (cut short) due to the incorporation of a "stop" signal into the DNA sequence. This results in translation being stopped before the amino acid sequence of the protein is completed.

MATERIALS

genetic code chart (see page 7 of this handout or a biology textbook)
blue, red, and green colored pencils



The Making of the Fittest: Natural Selection and Adaptation

PROCEDURE

1. Watch the short film *The Making of the Fittest: Natural Selection and Adaptation*.
2. Using the DNA nucleotide sequence in the gene tables (page 3), determine the complementary messenger RNA (mRNA) sequence for the portion of the *Mc1r* gene provided. (Note: You are only transcribing a small portion of the DNA sequence for this protein. The actual gene contains 951 base pairs.) The numbers above some of the columns indicate amino acid positions in the protein sequence. Also indicated is whether the sequence codes for an extracellular, intracellular, or transmembrane part of the protein.
3. Using the mRNA sequence determined in Step 2, determine the resulting amino acid sequence of the MC1R protein. (Note: This is only a portion of the 317 amino acids in the entire protein. The numbers above some of the columns in the tables indicate amino acid positions in the protein sequence.) You may use the genetic code chart provided in your textbook or the one on page 7 of this handout.
4. There are five mutations in the dark-color *Mc1r* mutant gene. Compare the DNA sequence of the wild-type *Mc1r* gene with the DNA sequence of the mutant *Mc1r* gene. **Indicate the locations of the five mutations by circling the five single DNA nucleotides that are mutated in the mutant *Mc1r* gene table.**
5. From the introduction, determine whether each of these mutations is a **silent, missense, or nonsense mutation.**
 - a. Using the mutant *Mc1r* gene data, shade in the columns (including DNA, mRNA, and amino acid) in the mutant table that contain a **silent** mutation. Use a **blue** colored pencil to do this.
 - b. Likewise, use a **red** colored pencil to shade in the columns that contain a **missense** mutation.
 - c. Shade any columns that contain **nonsense** mutations by using a **green** colored pencil.
6. Answer the questions following the gene tables.



The Making of the Fittest: Natural Selection and Adaptation

LESSON
STUDENT HANDOUT

GENE TABLES

WILD-TYPE MC1R GENE (LIGHT-COLORED COAT PHENOTYPE)

015											Extracellular Domain I
DNA	TTG	AGG	TGG	GCG	TGT	CCG	CAA	GGA	GTG	GAG	
mRNA											
Amino Acid											

MUTANT MC1R GENE (DARK-COLORED COAT PHENOTYPE)

015											Extracellular Domain I
DNA	TTG	AGG	TGG	ACG	TGT	CCG	CAA	GGA	GTG	GAG	
mRNA											
Amino Acid											

WILD-TYPE MC1R GENE (LIGHT-COLORED COAT PHENOTYPE)

105											Extracellular Domain III
DNA	CGG	GAC	CGG	TGG	GCC	CAC	TGA	CAC	CAT	GTC	
mRNA											
Amino Acid											

MUTANT MC1R GENE (DARK-COLORED COAT PHENOTYPE)

105											Extracellular Domain III
DNA	CGG	GAC	CGG	TGG	ACC	CAC	TGA	CAC	CAT	GTC	
mRNA											
Amino Acid											

WILD-TYPE MC1R GENE (LIGHT-COLORED COAT PHENOTYPE)

154											Extracellular Domain I
DNA	TCA	TAA	CAC	TGT	GAC	GGG	GCC	CGA	GCC	ACC	
mRNA											
Amino Acid											

MUTANT MC1R GENE (DARK-COLORED COAT PHENOTYPE)

154											Intracellular Domain I
DNA	TCA	TAA	CAC	TGT	GAC	GGG	ACC	CGA	GCC	ACC	
mRNA											
Amino Acid											



The Making of the Fittest: Natural Selection and Adaptation

**LESSON
STUDENT HANDOUT**

WILD-TYPE MC1R GENE (LIGHT-COLORED COAT PHENOTYPE)

DNA	CAC	GTG	TAC	GAA	CGT	Transmembrane V
mRNA						
Amino Acid						

DNA	CAC	GTG	TAC	GAA	CGT	Transmembrane V
mRNA						
Amino Acid						

DNA	CAC	GTG	TAC	GAA	CGT	Transmembrane V
mRNA						
Amino Acid						

DNA	CAC	GTG	TAC	GAA	CGT	Transmembrane V
mRNA						
Amino Acid						

DNA	CAC	GTG	TAC	GAA	CGT	Transmembrane V
mRNA						
Amino Acid						

DNA	CAC	GTG	TAC	GAA	CGT	Transmembrane V
mRNA						
Amino Acid						

DNA	CAC	GTG	TAC	GAA	CGT	Transmembrane V
mRNA						
Amino Acid						

DNA	CAC	GTG	TAC	GAA	CGT	Transmembrane V
mRNA						
Amino Acid						

DNA	CAC	GTG	TAC	GAA	CGT	Transmembrane V
mRNA						
Amino Acid						

DNA	CAC	GTG	TAC	GAA	CGT	Transmembrane V
mRNA						
Amino Acid						

DNA	CAC	GTG	TAC	GAA	CGT	Transmembrane V
mRNA						
Amino Acid						

DNA	CAC	GTG	TAC	GAA	CGT	Transmembrane V
mRNA						
Amino Acid						

DNA	CAC	GTG	TAC	GAA	CGT	Transmembrane V
mRNA						
Amino Acid						

DNA	CAC	GTG	TAC	GAA	CGT	Transmembrane V
mRNA						
Amino Acid						

DNA	CAC	GTG	TAC	GAA	CGT	Transmembrane V
mRNA						
Amino Acid						

DNA	CAC	GTG	TAC	GAA	CGT	Transmembrane V
mRNA						
Amino Acid						

DNA	CAC	GTG	TAC	GAA	CGT	Transmembrane V
mRNA						
Amino Acid						

DNA	CAC	GTG	TAC	GAA	CGT	Transmembrane V
mRNA						
Amino Acid						

DNA	CAC	GTG	TAC	GAA	CGT	Transmembrane V
mRNA						
Amino Acid						

DNA	CAC	GTG	TAC	GAA	CGT	Transmembrane V
mRNA						
Amino Acid						

DNA	CAC	GTG	TAC	GAA	CGT	Transmembrane V
mRNA						
Amino Acid						

DNA	CAC	GTG	TAC	GAA	CGT	Transmembrane V
mRNA						
Amino Acid						

DNA	CAC	GTG	TAC	GAA	CGT	Transmembrane V
mRNA						
Amino Acid						

DNA	CAC	GTG	TAC	GAA	CGT	Transmembrane V
mRNA						
Amino Acid						

DNA	CAC	GTG	TAC	GAA	CGT	Transmembrane V
mRNA						
Amino Acid						

DNA	CAC	GTG	TAC	GAA	CGT	Transmembrane V
mRNA						
Amino Acid						

DNA	CAC	GTG	TAC	GAA	CGT	Transmembrane V
mRNA						
Amino Acid						

DNA	CAC	GTG	TAC	GAA	CGT	Transmembrane V
mRNA						
Amino Acid						

DNA	CAC	GTG	TAC	GAA	CGT	Transmembrane V
mRNA						
Amino Acid						

DNA	CAC	GTG	TAC	GAA	CGT	Transmembrane V
mRNA						
Amino Acid						

DNA	CAC	GTG	TAC	GAA	CGT	Transmembrane V
mRNA						
Amino Acid						

DNA	CAC	GTG	TAC	GAA	CGT	Transmembrane V
mRNA						
Amino Acid						

DNA	CAC	GTG	TAC	GAA	CGT	Transmembrane V
mRNA						
Amino Acid						

DNA	CAC	GTG	TAC	GAA	CGT	Transmembrane V
mRNA						
Amino Acid						

DNA	CAC	GTG	TAC	GAA	CGT	Transmembrane V
mRNA						
Amino Acid						

DNA	CAC	GTG	TAC	GAA	CGT	Transmembrane V
mRNA						
Amino Acid						

DNA	CAC	GTG	TAC	GAA	CGT	Transmembrane V
mRNA						
Amino Acid						

DNA	CAC	GTG	TAC	GAA	CGT	Transmembrane V
mRNA						
Amino Acid						

DNA	CAC	GTG	TAC	GAA	CGT	Transmembrane V
mRNA						
Amino Acid						

DNA	CAC	GTG	TAC	GAA	CGT	Transmembrane V
mRNA						



The Making of the Fittest: Natural Selection and Adaptation

QUESTIONS

1. Using the amino acid numbers provided above the first and last column of each table, list the **locations** of the five amino acids that contain a mutation.

2. Of the five mutations you identified in the *Mc1r* gene, how many are the following:

____ substitutions ____ insertions ____ deletions (Enter a number on each line.)

3. Of the five mutations you identified in the *Mc1r* gene, how many are the following:

____ silent ____ missense ____ nonsense (Enter a number on each line.)

4.

a. Which four amino acid locations (see Question 1 above) contain the missense mutations?

b. Explain the link between DNA sequence and protein structure and function.

5. Using the information on mutations in the introduction and your knowledge of proteins, develop a hypothesis to explain how the changes in the MC1R protein's amino acid sequence might affect its function.

6. Using the information on the *Mc1r* gene in the introduction and your knowledge of proteins, develop a hypothesis to explain how the change in MC1R protein function might directly affect a rock pocket mouse's coat color. Be specific and consider both the light-colored and dark-colored phenotypes.

8. Explain why the mutation at amino acid location 211 is not as significant as the other four mutations.

9. Mutations are a source of genetic variation. In the film, Dr. Sean Carroll says that mutations occur randomly. What does this mean?



The Making of the Fittest: Natural Selection and Adaptation



- 10.** It is a common misconception that “all mutations are bad.” Use the example of rock pocket mice to explain why this statement is not true. In your answer, explain how the dark coat-color mutation can be an advantage to some mice and a disadvantage to others.

- 11.** Use your understanding of evolution and the information in the film to explain how the dark-colored mutation came to be so common in some populations of rock pocket mice. Be specific.
