

1 A cell must continuously sort useful materials from its waste products and then remove or recycle the waste. One way a cell does this is to first put **Ubiquitin** tags on proteins that are no longer needed. These tags, which are small polypeptide chains 76 amino acids long, are called **ubiquitin**.

2 **Ubiquitin** tags direct proteins to compartments of the cell called **proteasomes**. Within each proteasome, ubiquitin-tagged proteins are deconstructed into their component amino acids. These pieces can be repurposed to build new proteins.

3 Sometimes organs stop processing waste effectively, allowing the waste to pile up and contributing to many diseases. Alzheimer's disease, for example, includes piles of waste proteins in the brain called **tangles**, while Parkinson's disease involves the accumulation of recyclable material into **Lewy bodies**. **Huntington's bodies** are piles of waste proteins common in liver disease. These piles, known generically as **inclusions**, can be identified under the microscope because they are covered in tiny ubiquitin tags.

4 A problem in the breakdown of ubiquitin-tagged proteins may be one cause of **spinocerebellar atrophy type 1**. In this disease, a protein called **ataxin-1** has an **abnormal amino acid**, which makes it able to attach to the ubiquitin **tags**. Scientists believe that the mutated protein cannot be chopped up by the proteasome because of its shape, so ubiquitin-tagged ataxin-1 starts to **accumulate**, which leads to **cell death**. The cells most affected by the genetic mutation are neurons in a part of the brain associated with **motor control** and the **spinal cord**. So, a person with mutant ataxin-1 progressively loses muscle control as these cells die.

Why do individuals with spinocerebellar atrophy experience loss of **muscle control**, even though the ataxin-1 protein builds up only in the brain and spinal cord?

- A. Inclusions can build up in all body systems and can cause debilitating effects.
- B. Genetic mutations in the ataxin-1 protein cause molecular proteins to change shape.
- C. When the ubiquitin tagging system does not work effectively, cellular waste can build up in the muscles.
- D. The neurons affected by the ataxin-1 protein buildup are in a region of the brain that manages muscle control.

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