Teaching meiosis with the DNA triangle framework: A classroom activity that changes how students think about chromosomes

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Abstract
Many biology students struggle to learn about the process of meiosis and have particular difficulty understanding the molecular basis of crossing over and the importance of homologous pairing for proper segregation. To help students overcome these challenges, we designed an activity that uses a newly developed Chromosome Connections Kit® from 3-D Molecular Designs to allow learners to explore meiosis at the molecular level. We took a backwards design approach in constructing an effective classroom activity. We developed evidence-based learning objectives and designed a crossing over activity that targets students' misconceptions and key concepts about meiosis. Assessment questions were designed based on the learning objectives and common student misconceptions. The activity consists of three parts: an interactive introductory video, a model-based activity, and reflection questions. The activity was first beta-tested with a small number of students and revised based on feedback. The revised activity was deployed in a mid-level Cell and Molecular Biology course. Analysis of pre-/post-assessment data from students who completed the activity (n = 83) showed strong learning gains on concepts related to ploidy, homology, segregation, and the mechanism and purpose of crossing over. Additionally, students who participated in the activity outperformed non-participants on a Genetics assessment about meiosis the following semester.

KEYWORDS
active learning, assessment of educational activities, molecular biology, student conceptual and reasoning difficulties

1 INTRODUCTION

Getting students to understand the process of meiosis is a common struggle for undergraduate biology instructors. College students are often bored or frustrated when presented with information about meiosis in their classrooms because they (mistakenly) think they already have solid knowledge about this subject. Since typical instruction
about meiosis often begins in middle school, is repeated in high school, and then again in college Introductory Biology, and yet again in Cell Biology and Genetics courses, it is not surprising that students are tired of reading, seeing, and/or hearing about this process. Contrary to what many students may think about their own mastery of the subject, the biology education literature provides ample evidence that students hold numerous misconceptions, incomplete and/or incorrect ideas about this process.\textsuperscript{1–9} When students do not have deep knowledge about the underlying reasons and concepts linked with meiosis, it hinders their ability to master future concepts in genetics and evolution. We have focused much of our research on understanding how learners conceptualize meiosis and how their mental models of this process differ from that of experts.\textsuperscript{10,11} One of the most important differences between experts and novices is that experts rely on molecular level knowledge about DNA when they describe certain concepts related to meiosis, learners do not.

The underlying processes that drive meiosis (i.e., meiotic recombination) are not directly observable, so instructional materials usually rely on simplified drawings to help learners visualize what is happening within a cell. Biology experts, who have both content knowledge and visual literacy skills, can create and decipher visual representations to communicate, ask, and answer questions.\textsuperscript{12–14} Learners, on the other hand, do not have the experience or the visual literacy skills of experts\textsuperscript{15–18} and may not learn as much from static textbook images as instructors would hope. Bolstered by previous research findings about the deficiencies of traditional textbooks in illustrating the molecular level of meiotic processes,\textsuperscript{11,19} we posit that students struggle to understand meiosis because they see (when looking at textbook images as well as when picturing the process in their minds) chromosomes in a two-dimensional way—usually as red or blue blob-like structures. Chromosomes, however, are incredibly complex molecules of DNA and protein. DNA encodes genetic information in both a concrete (sequence of nucleotide bases) and an abstract way (regulatory regions of genes influence expression, which can influence phenotype). The DNA helix and nucleotide bases cannot be visualized directly and yet, the nucleotide sequence promotes or represses molecular level interactions that drive expression and chromosome behavior. Macroscopic level chromosome structures also present problems for learners; experts understand why and how chromosomes can be one or two-DNA molecules, but students get confused as to what counts as a chromosome and where the structures originated from. To be proficient at understanding meiosis one must integrate three levels of DNA knowledge (molecular, chromosomal, and informational) into their mental model. Experts can do this readily, but students cannot.

The DNA Triangle framework is an established model that serves to reconcile and integrate these three different levels of DNA into one cogent mental model.\textsuperscript{11} The corners of the triangle represent each of the three levels of DNA that must be conceptualized when teaching or learning about a process such as meiosis. The chromosomal (C) level describes the structure of chromosomes (with and without sister chromatids), identification of chromosomes by banding pattern and centromere location, representations of chromatin packing, and counting chromosomes. The informational (I) level describes how DNA encodes genetic information, such as genes or alleles, protein-coding regions, or regulatory information. Lastly, the molecular (M) level describes the chemistry and nucleotide sequence of DNA. Unfortunately, college-level biology textbooks typically describe (by the text as well as the visual representations) important concepts like ploidy, homology, and segregation at only the chromosomal level.\textsuperscript{11,19} Typical models or activities designed to help students learn meiosis often involve manipulatives such as pool noodles, pop-beads, or socks.\textsuperscript{7,20,21} While a hands-on “active-engagement” approach is certainly better than a straight lecture,\textsuperscript{22–24} most models fall short because they mainly focus on the chromosomal level and only hint at the informational level. A classic example is a pop-bead kit in which students build pairs of “homologous chromosomes” with red or blue beads. Color (blue or red beads) represent origin (paternal or maternal) but the lack of nucleotide sequence and molecular level structure encourages students to only focus on superficial aspects such as length and size of “homologous” chromosomes. Beads often represent different genes with different colors representing different alleles, but this model focuses mainly on the chromosome structure, which is highly problematic for several reasons. The scale is completely nonsensical (if there are 12 beads on the chromosomes, do chromosomes only contain 12 genes?) and alleles are versions of genes, which are sequences of nucleotides. The beads look more like a protein than they do DNA. In addition, all of the beads on a chromosome are identical, which makes it confusing as to what an “allele” actually is. Lastly, students are encouraged to exchange red and blue beads to mimic the results of crossing over, but there is no instructional cue or information presented on how that process actually works.

Out of all the parts of meiosis, the process of crossing over for proper chromosome segregation is likely the most mysterious to students.\textsuperscript{7,25} Most textbooks illustrate the process of crossing over (meiotic recombination) and resolution very poorly. Many textbook illustrations simply show a pair of homologous chromosomes, one red and one blue, lined up side-by-side (no physical interaction) and do not show individual strands of DNA nor
how they invade and combine with a complementary strand on a non-sister chromatid. Hand-held models such as pop-bead chromosomes show crossing over by exchanging red and blue beads on non-sister chromatids; this model serves to strengthen a common misconception or misunderstanding about crossing over, namely that chromatid pieces simply break off and exchange places.

The process of crossing over (meiotic recombination) is essential for proper chromosome alignment and segregation, and failure to maintain the integrity of this process has severe consequences for a cell. The basic mechanisms that drive meiotic recombination arise from the double-strand break repair pathway, a process that is of critical importance in numerous molecular and genetics topics. Yet the underlying molecular nature of the crossing-over process is rarely described or illustrated in undergraduate biology textbooks, especially introductory levels. Molecular-level interactions that drive cellular processes cannot be directly visualized and, thus, may be “black boxes” for the novice student trying to comprehend these processes. We hypothesized that revealing the molecular level would help students understand how and why homologous DNA sequence drives the process of crossing over and why it is essential for proper segregation.

With the advancement of three-dimensional (3D) printing and other educational technologies, it is now possible to bring physical models of biomolecules (DNA, RNA, and proteins) into the undergraduate classroom. Physical models provide opportunities for students to compare and revise their own mental models of scientific phenomena and have been shown to produce increased learning gains compared to other active learning strategies. Thus, we set out to design an activity to explicitly reveal the molecular level to help students visualize homology and the process of crossing over. Following evidence-based practice, we wanted the activity to require the students to construct their own understanding of the process through the manipulation of physical models.

2 DEVELOPMENT OF THE MODEL

The Center for BioMolecular Modeling (CBM) is a grants-funded instructional materials development laboratory that focuses on developing tactile materials to explore challenging concepts at the molecular level. The CBM offers a number of professional development opportunities for high school and college educators who provide input on which concepts are difficult for students to grasp. As part of an NIH-SEPA grant focusing on genome editing, educator colleagues identified mitosis and meiosis as challenging topics to teach. Many existing tools focus on the movement of chromosomes during cell division (a cytological view), and some make the connection between alleles and chromosomes. But none of the cell division models made the connection between chromosomes and nucleotides (a molecular perspective). Discussions with LKW and DLN regarding student misconceptions, along with the DNA Triangle framework guided the design of foam chromosome models that allow for a “zoom in” on the chromosome to reveal that it contains double-stranded DNA (see Figure 1). The modularity of the foam models allows for the exploration of cell division (mitosis and meiosis) at both the cytological and molecular scales, thereby solidifying the connection between DNA and chromosomes. The kits went through several iterations of field-testing in high school and college classrooms, and educator feedback guided final model designs. Red and blue (signifying maternal and paternal origin) foam pieces fit together like puzzle pieces, to build chromosomes at the macroscopic/chromosomal level and nucleotide pieces (also red or blue) to allow learners to appreciate the molecular structure and sequence of chromosomes. The nucleotide sequences embedded within the chromosome allow for the exploration of advanced topics, including the molecular mechanism of crossing over and the creation of a Holliday junction. They can also be used to explore chromosomal aberrations and recombination repair of damaged DNA. These kits are now produced by and available from 3D Molecular Designs as the Chromosome Connections Kit© and are also available for loan from the MSOE Model Lending Library (https://cbm.msoe.edu/lendingLibrary).

3 METHODS

Following the three main steps in backwards design—identify desired results, determine acceptable evidence, plan learning experiences, and instruction—we developed the activity and assessment materials to ensure that the assessment, activities, and learning objectives (LOs) were in alignment. The process was iterative. For example, as we crafted the LOs we also considered assessment questions and what, specifically, students would do with the models in class.

3.1 Development of learning objectives

Using the literature and our own teaching experiences we drafted a set of LOs. As we developed the assessment and activities, we continued to refine the LOs for improved clarity and readability. Table 1 lists the LOs,
FIGURE 1 The crossing over activity is based on the Chromosome Kit® from 3D molecular designs. It contains foam pieces that interlock and can be used to build chromosomes. (a) Red and blue pieces can be used to represent maternal and paternal chromosomes. Inserts allow for demonstration of “zooming in” to the sequence level, where individual nucleotides can be added. (b) The three types of parts are put together to show the sequence similarity of homologous chromosomes.

TABLE 1 Learning objectives were developed based on research about student misconceptions and naïve ideas about meiosis

| Learning objective | Students will be able to: | Examples of common novice ideas/misconceptions related to LO |
|--------------------|---------------------------|----------------------------------------------------------|
| LO1                | Correctly predict chromosome number at any stage of meiosis. | Students incorrectly count chromatids as chromosomes, leading to incorrect conclusions about when chromosome number is changed.6 |
| LO2                | Correctly count chromosomes and chromatids at all stages of meiosis. | Chromosome structure (replicated vs. unreplicated) determines chromosome number.1,3 One replicated chromosome with two chromatids are comprised of one maternal and one paternal chromosome.34 |
| LO3                | Identify the basis of homology. | Homologous is based on size and shape of chromosomes (only). Homologous chromosomes carry similar genetic information but cannot link the concept of “genetic information” to the nucleotide sequence.11,25 |
| LO4                | Describe the process of crossing over at the molecular level. | Homologous chromosomes pair because of chaperone proteins (not complementary DNA sequence).10 Crossing over involves pieces of chromatids changing place, but the mechanism is unknown.11 |
| LO5                | Correctly outline the major steps of meiosis. | Students do not remember that DNA replication precedes meiosis. If there is an odd number of homologous pairs students add or subtract chromosomes to make the final gametes come out “correct.”10 |
| LO6                | Describe possible outcomes of crossing over. | New to this research (related idea is that students struggle with recombination). |
| LO7                | Explain why crossing over is necessary for homologous chromosomes to segregate properly. | Crossing over is “not necessary” for the process of meiosis to occur correctly.10,25 |
| LO8                | Apply their knowledge of ploidy in distinguishing the types of cells that result after both stages of meiosis. | Cells become haploid only at the end of meiosis II.7,11 |
| LO9                | Correctly identify where crossing over occurs in meiosis. | Students do not always incorporate crossing over into mental models of meiosis occurs10 and/or are confused where in the process crossing over occurs.6 |
references, and examples of common novice idea/misconceptions that helped informed each one.

### 3.2 Development of assessment

Guided by the LOs a series of multiple select assessment questions (i.e., “select all that apply”) were developed. Four answer choices were created for each question stem and between 1 and 4 answer choices were correct for each question. Questions and answer choices (both correct and incorrect) were mapped back to the LOs and questions were revised to ensure coverage of all LOs. With the exception of LO4, all objectives aligned with both correct and incorrect answer choices. The assessment is included as a Data S1.

### 3.3 Development of the activity

D.L.N. and L.K.W. collaborated with the team at the CBM on early model prototypes. The starting point for the model was the paper chromosomes described in Reference 25, which was then transitioned to a more manipulable and durable foam structure. The basic concept of the design was to create a model that allowed for users to zoom in (to M level) and out (to C level) and to incorporate allelic differences (I level) in order to bring all parts of the DNA Triangle together. This allows users to manipulate chromosomes to show large-scale movements (C level) but also to get insight into how similar homologues are (M level), how the process of crossing over works (M level), and the genetic outcomes (I level). Once the model was fully developed, the research team worked with the model to develop LOs and conceptualize how students would move the model pieces around to accomplish the LOs. Step by step instructions were developed and revised. The activity was given to four beta-testers, who explained their thinking while following instructions and manipulating the models. Their experiences revealed confusing steps/language and the fact that the activity was too long for one in-class setting. In the second major revision, the activity was divided into three parts, one of which included a video. Table 2 provides a

| Phase                          | Synopsis                                                                 | Key concepts     | Levels |
|--------------------------------|--------------------------------------------------------------------------|------------------|--------|
| Pre-activity assessment        | Eleven item multiple select assessments. Given in an online format.       | N/A              | N/A    |
| Part I: In-class model-based activity (30 min) | Students were placed in small groups (4–5 students), given the red and blue foam pieces from the 3DMD Chromosome Kit and an activity sheet. Students were challenged to build chromosomes and model their movement through the major steps of meiosis of a diploid cell with three pairs of chromosomes. The instructor and Learning Assistant walked around, provided feedback, and questioned students about their models. | Ploidy           | C, I   |
| Part II: Video (10 min)        | Students watched a video reviewing chromosome counting and ploidy. The narrator reintroduced the foam chromosome pieces and probed students to think about chromosome number, structure, ploidy, and homology. Students watched the video as a pre-class homework assignment in preparation for class. The video contained five multiple choice questions to keep students engaged during the video. | Ploidy, homology | C, I   |
| Part III: In-class model-based activity (60 min) | Students were placed in small groups (4–5 students), given the 3DMD Chromosome Kit and an activity sheet. Students built homologous chromosomes using foam nucleotide pieces (red and blue) and modeled the processing of crossing over and recombination by building a Holliday junction and resolving the structure. The instructor and Learning Assistant walked around, provided feedback, and questioned students about their models. | Homology, segregation | I, M C, M |
| Post-activity assessment       | Eleven item multiple select assessment (same as pretest) given in paper format. Questions were embedded in the final course exam. | N/A              | N/A    |

Abbreviations: C, chromosomal; I, informational; M, molecular.
synopsis of the activity (all three parts) and how they align with levels of DNA as described in the DNA Triangle framework. Figure 2 shows students constructing a Holliday junction with the model. Assessments and the activity are available as Data S1.

3.4 Implementation of activity

The crossing over activity was tested in a second-year Cell and Molecular Biology majors-level course at a large, private institution in the Northeast. Students enrolled in this course were biology, biochemistry, or other biology-related majors or were pursuing their minor or concentration in biology. All completed a year of introductory biology before enrollment in this course. There were three identical lecture sections of this course taught by two different instructors (author L.K.W. was one of the instructors but administered the model-based activity in all three sections for this study). One week prior to the implementation of Part I of the activity, students completed the multiple select pre-assessment given in an online format. Students were awarded homework points for their participation and “best effort” but scores were not shared nor were questions discussed. During class, students completed Part I of the crossing over activity. After class was complete, students watched a short review video on their own (Part II of the activity) before coming to the subsequent class to complete Part III of the activity. Of 93 students enrolled in the course, 83 completed the pretest, posttest, and all parts of the activity. Figure 3 provides an overview of the student populations included in this study.

3.5 Analysis of assessment data

Pre and posttests from students who completed the crossing over activity (n = 83) were analyzed two ways. We calculated pre- and post-scores for each student based on whole question and also partial responses. This dual scoring strategy (whole question and partial or “fractional”) has been utilized in other studies such as References 35, 36. For whole question analysis, students got credit for each question they answered perfectly correct (selecting all correct choices and not selecting any incorrect choices). The percentage of each question students answered correctly was calculated for partial answer scores. We also analyzed assessment data based on LOs. Table 3 illustrates the alignment of LOs with correct and incorrect answer choices. Normalized learning gains for each LO were calculated using (post-pre)/(1-pre). We also calculated the percent change in selection of correct answers or incorrect answers on pretest compared with posttest for each LO.

To investigate longer-term impacts on learning, we analyzed student performance on a meiosis-related question that appeared on a Genetics in-class exam 5 weeks into the semester, after all students had reviewed meiosis. Genetics is a course that students take after Cell and
Molecular Biology, although not always in the next consecutive semester. Thus, about half of the students enrolled in Genetics had used the model when it was first introduced the previous semester and half had taken the course earlier, without the model. Thirty-six students were enrolled in the course. The instructor of the Genetics course (author D.L.N.) did not have prior knowledge as to which students had used the model previously. In the exam question, students were presented with illustrations of various cells containing replicated or unreplicated chromosomes and asked to determine the total number of chromosomes, number of homologous pairs, and number of double-stranded DNA molecules, and to decide whether each was haploid or diploid and whether or not replication had already taken place. They were also asked to place alleles on one diagram to indicate a triple heterozygote where one pair was linked, and to predict the gametes produced by that individual. After the exams were all graded, co-author L.K.W. looked at the class-list and sorted the 36 students enrolled in the Genetics class. Nineteen of the students were not exposed to the model ($N = 19$, “controls”) and 15 students were exposed to the model ($N = 15$, “cases”). Two students in Genetics had partial experience with the model and were excluded from the analysis.

![Molecular Biology and Genetics Class](image)

**Figure 3** Data for this study was drawn from two populations. First, pre-/post-scores were calculated from 83 students enrolled in a cell and molecular biology class, who completed all parts of the model-based activity and pre/posttests. A subset of the 83 students enrolled the following semester in a Genetics class. The second analysis compared the 15 students exposed to the model to 19 students who had not been exposed to the model. Two students in Genetics had partial experience with the model and were excluded from the analysis.

| Learning objective | Students will be able to: | Correct | Incorrect | Average normalized learning gain |
|--------------------|---------------------------|---------|-----------|---------------------------------|
| LO1                | Correctly predict chromosome number at any stage of meiosis. | 1b, 1d, 2a, 2b | 1a, 1c, 9c | 0.2875 |
| LO2                | Correctly count chromosomes and chromatids at all stages of meiosis. | 4a, 4b, 4c, 4d | 2c, 3b | 0.3309 |
| LO3                | Identify the basis of homology. | 5c | 5a, 5b, 5d | 0.1172 |
| LO4                | Describe the process of crossing over at the molecular level. | 6a, 6b, 6c, 6d | | 0.4914 |
| LO5                | Correctly outline three major steps of meiosis. | 3a, 8c, 9d | 3c, 3d | 0.4422 |
| LO6                | Describe possible outcomes of crossing over. | 8a, 8b | 8d | 0.3788 |
| LO7                | Explain why crossing over is necessary for homologous chromosomes to segregate properly. | 7a, 7c, 7d | 7b, 10d | 0.4860 |
| LO8                | Apply their knowledge on ploidy in distinguishing the types of cells that result after both stages of meiosis. | 9a | 2d, 9b | 0.3400 |
| LO9                | Correctly identify where crossing over occurs in meiosis. | 10c | 10a, 10b | 0.4691 |

**Table 3** Alignment of learning objectives with correct and incorrect answer choices from the multiple select assessment

Note: Average normalized learning gains for each learning objective were also calculated.
therefore, were not included in the analysis because they had partial exposure but did not complete the entire activity as intended. Median scores and effect size for each group were calculated.

4 | RESULTS

4.1 | Pre and posttesting reveals learning

Analysis of the pre and post-assessment scores strongly suggests students learned as a result of using the crossing-over model. The median score on the pretest assessment was a 61.63%, which improved to a 72.72% on the posttest (Figure 4).

The multiple select nature of the assessment improves the validity of the assessment, as students cannot use typical “game playing” strategies to identify just one correct answer choice. Students must evaluate each answer choice as being correct or incorrect which also increases the difficulty of the assessment. The multiple select formats allow students to be both partially incorrect and partially correct at once, which offers instructors and researchers a more complete picture of student understanding compared to a standard forced-choice assessment.36,37 We found that students make learning gains on all LOs (Table 3), with particularly strong gains on LO4, LO7, and LO9. Students made the smallest gains on LO3, to “Identify the basis of homology.” Deeper analysis into the responses reveals the vast majority of students (89%) correctly recognize choice 5c that “Both chromosomes of the pair have similar DNA sequence” on the posttest but they do not easily stop choosing 5a and 5b, which are “both chromosomes of the pair have the same alleles” and “both chromosomes of the pair are similar in length”. While choices 5a and 5b are not untrue statements, they are not the underlying basis of homology.

As the pre-/post-assessment followed a multiple select format we were able to calculate the percent change in selection of correct and incorrect answers on post compared to pre, aligned by each LO. As demonstrated by Figure 5, students selected more correct choices and fewer incorrect choices after engaging in the crossing over activity. The pattern varied somewhat by LO, but students improved on all objectives in the post-assessment analysis. This outcome is especially pronounced in LO7 (Students will be able to explain why crossing over is necessary for homologous chromosomes to segregate properly), which was a major focus of the activity. Since there were no incorrect answer options for LO4, only the increase in correct responses could be calculated.

4.2 | Long-term effects

To assess any long-term impacts of the crossing over activity, we analyzed data from a genetics exam question given the following semester. The Cell and Molecular course is a prerequisite for Genetics, but students do not always take Genetics immediately after Cell and Molecular Biology. Of the students enrolled in the Genetics course, 15 students did the crossing over activity and 19 students did not (they had completed the Cell and Molecular Biology course in a previous term). As illustrated by Figure 6, students who had previously used the model outperformed students who did not. Students who

\[ \begin{align*}
\text{Score} & \quad 0 \quad 10 \quad 20 \quad 30 \quad 40 \quad 50 \\
\text{Pre} & \quad (\text{N} = 83) \\
\text{Post} & \quad (\text{N} = 83)
\end{align*} \]

**FIGURE 4** Students (N = 83) made significant improvements on their posttest scores, suggesting they gained knowledge on the learning objectives aligned with the activity

\[ P < 0.00001 \]

**FIGURE 5** Students (N = 83) chose more correct options and fewer incorrect options on the posttest compared to the pretest for every learning objective (note that there were no incorrect options offered for LO4). The bar graph shows the change in the percentage of correct answers (positive change, more correct answers selected) and incorrect answers (negative change, fewer wrong answers selected) in the posttest compared with the pretest.
used the model (cases) had a mean score of eight (of 10) points while students who did not (controls) had a mean score of 6.1. Effect size was calculated at 0.9939, which shows an extremely strong relationship between exposure to the model and exam question performance.38

5 | DISCUSSION

Visualizing the molecular level of DNA and seeing its relationship to the other levels are key to understanding the behavior of chromosomes (and the genetic consequences) during the process of meiosis. Traditional classroom activities and textbook illustrations of meiosis focus almost exclusively on the chromosomal and informational level and fail to incorporate the molecular level.19,39 Students are exposed to these same sets of teaching materials time and time again and are rarely challenged to think about the underlying molecular processes that drive chromosomal behavior. We designed our novel set of materials using the DNA Triangle framework as a guide to help students “see” the underlying molecular structure (nucleotide sequence) of homologous chromosomes and explore how sequence similarity drives the process of crossing over. The dynamic, 3D model also allows students to build the physical interaction of homologous chromosomes during crossing-over. This physical interaction is essential for proper chromosome alignment and eventual chromosome segregation; a fact that is not made clear to them with most other teaching approaches. Data analysis of pre-/post-assessment strongly suggests this crossing-over activity is an effective tool for learning. Because we did not have a control group of students who did not complete the activity but also completed pre/posttesting, we cannot rule out that students may have learned these concepts on their own and/or during some other part of the class. However, the quasi-random study of a subset of students who went on to a Genetics course compared to a control group who never experienced the activity supports our argument that the model-based activity was the element that improved learning.

Although the Chromosome Kit is available to borrow from the MSOE CBM Lending library (https://cbm.msoe.edu/lendingLibrary/), not all instructors will want to or will be able to use the kit and our three-part activity in their classrooms. Regardless, we feel there are still valuable ideas that instructors can use from our study, as discussed below.

5.1 | Implications for teaching

Our current study illustrates how important it is to let students “see” the molecular level of DNA; especially when learning about molecular processes that depend on the underlying nucleotide sequence. For instance, the concept of homology is rooted in the molecular level so we strongly suggest instructors use something other than standard textbook-like illustrations that depict homologous chromosomes as cartoonish red and blue chromosomes.19 Such illustrations direct students to focus on the color and size and shape of the chromosomes rather than the underlying sequence similarity. As a result, many students are unaware of the molecular basis of homology. The partial DNA sequences we provided for a maternal and paternal homologous pair (Data S1) may be a good way to illustrate the concept without using the kit-based model.

The process of crossing over is both familiar but also confusing to the typical biology student. Most students will remember that crossing over happens in meiosis, but the molecular details of the process are missing from students’ mental models.11 Most textbooks give little detail about the process and, instead, show the product of crossing over with cartoonish red and blue chromosomes that have “swapped” pieces. We argue that these figures are misleading, as they seem to show a piece from a “red” chromosome breaking off and fusing with a “blue” chromosome. The process of crossing over becomes much less mysterious to students when they can see a single paternal strand base-pair with a complementary maternal strand. If the kit-based model that we described here is not possible to bring into class, a paper-based model may be a useful alternative for instructors.25

Explicitly bringing in the molecular level of DNA does not begin and end with the process of meiosis. Many
processes are facilitated by complementary base-pairing such as DNA repair pathways (homologous recombination after double-strand break), hairpin loop-mediated transcriptional termination, and even tRNA:mRNA binding during translation. Molecular biology experts can “see” (in their mental models) nucleotides and the interactions that depend on precise nucleotide sequences, but learners (novices) cannot. Thus, when teaching topics in Molecular Biology we encourage instructors to find ways to make the molecular level visible to their students.

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REFERENCES
1. Kindfield ACH. Confusing chromosome number and structure: a common student error. J Biol Educ. 1991;25:193–200.
2. Kindfield ACH. Understanding a basic biological process: expert and novice models of meiosis. Sci Educ. 1994;78:255–83.
3. Kalas P, O’Neill A, Pollack C, Birol G. Development of a meiosis concept inventory. CBE Life Sci Educ. 2013;12:655–64.
4. Brown CR. Some misconceptions in meiosis shown by students responding to an advanced level practical examination question in biology. J Biol Educ. 1990;24:182–6.
5. Stewart J, Dale M. High school students’ understanding of chromosome/gene behavior during meiosis. Sci Educ. 1989;73:501–21. https://doi.org/10.1002/sce.370730410
6. Dikmenli M. Misconceptions of cell division held by student teachers in biology: a drawing analysis. Sci Res Essay. 2010;5:235–47.
7. Wright LK, Newman DL. An interactive modeling lesson increases students’ understanding of ploidy during meiosis. Biochem Mol Biol Educ. 2011;39:344–51.
8. Ozcan T, Yildirim O, Ozgur S. Determining of the university freshmen students’ misconceptions and alternative conceptions about mitosis and meiosis. Procedia Soc Behav Sci. 2012;46:3677–80. https://www.sciencedirect.com/science/article/pii/S1877042812018629
9. Smith MK, Knight JK. Using the genetics concept assessment to document persistent conceptual difficulties in undergraduate genetics courses. Genetics. 2012;191:21–32.
10. Newman DL, Catavero C, Wright Students LK. Fail to transfer knowledge of chromosome structure to topics pertaining to cell division. CBE Life Sci Educ. 2012;11:425–56.
11. Wright LK, Catavero CM, Newman DL. The DNA triangle and its application to learning meiosis. Cell Biol Educ. 2017;16:1–14.
12. Towns MH, Raker JR, Becker N, Harle M, Sutcliffe J. The biochemistry tetrahedron and the development of the taxonomy of biochemistry external representations (TOBER). Chem Educ Res Pract. 2012;13:296–306. https://pubs.rsc.org/en/content/articlehtml/2012/rp/c2rp00014h
13. Trumbo Visual J. Literacy and science communication. Sci Commun. 1999;20:409–25.
14. Schönborn KJ, Anderson TR. The importance of visual literacy in the education of biochemists*. Biochem Mol Biol Educ. 2006;34:94–102.
15. Halverson KL, Friedrichsen P. In: Treagust DF, Tsui C-Y, editors. Multiple representations in biological education. Dordrecht: Springer Netherlands; 2013. p. 185–201. https://link.springer.com/book/10.1007/978-94-007-4192-8
16. Hinze SR, Rapp DN, Williamson VM, Shultz MJ, Deslongchamps G, Williamson KC. Beyond ball-and-stick: Students’ processing of novel STEM visualizations. Learn Instr. 2013;26:12–21. https://www.sciencedirect.com/science/article/pii/S0959475212001065
17. McCollum BM, Regier L, Leong J, Simpson S, Sterner S. The effects of Using touch-screen devices on Students’ molecular visualization and representational competence skills. J Chem Educ. 2014;91:1810–7.
18. Steiff M, Scopelitis S, Lira ME, Desutter Improving D. Representational competence with concrete models. Sci Educ. 2016;100:344–63.
19. Wright LK, Dy GE, Newman D. Undergraduate textbook representations of meiosis neglect essential elements. Am Biol Teach. 2020;82:296–305.
20. Locke J, McDermid Using HE. Pool noodles to teach mitosis and meiosis. Genetics. 2005;170:5–6. https://doi.org/10.1534/genetics.104.032060
21. Wells Simple RF. Chromosome models. Am Biol Teach. 1982;44:311–2.
22. Freeman S, Eddy SL, McDonough M, Smith MK, Okoroafor N, Jordt H, et al. Active learning increases student performance in science, engineering, and mathematics. PNAS. 2014;111:8410–5.
23. Theobald EJ, Hill MJ, Tran E, Agrawal S, Arroyo EN, Behling S, et al. Active learning narrows achievement gaps for underrepresented students in undergraduate science, technology, engineering, and math. PNAS. 2020;117:6476–83.
24. Wiggins BL, Eddy SL, Grunspan DZ, Crowe AJ. The ICAP active learning framework predicts the learning gains observed in intensely active classroom experiences. AERA Open. 2017;3:1–14. https://journals.sagepub.com/doi/pdf/10.1177/2332858417708567
25. Newman DL, Wright LK. Meiosis: a play in three acts, starring DNA sequence. CourseSource. 2017;4:1–9. https://doi.org/10.24918/cs.2017.9
26. Páques F, Haber Multiple JE. Pathways of recombination induced by double-strand breaks in Saccharomyces cerevisiae. Microbiol Mol Biol Rev. 1999;63:349–404.
27. Herman T, Morris J, Colton S, Batiza A, Patrick M, Franken M, et al. Tactile teaching: exploring protein structure/function using physical models*. Biochem Mol Biol Educ. 2006;34:247–54.
28. Cooper AK, Oliver-Hoyo Creating MT. 3D physical models to probe student understanding of macromolecular structure: Creating 3D physical models. Biochem Mol Biol Educ. 2017;45:491–500.
29. Babilonia-Rosa MA, Kuo HK, Oliver-Hoyo Using MT. 3D printed physical models to monitor knowledge integration in biochemistry. Chem Educ Res Pract. 2018;19:1199–215.
30. Terrell CR, Franzen MA, Herman T, Malapati S, Newman DL, Wright LK. Biochemistry Education: From Theory to Practice. American Chemical Society; 2019. p. 43–62.
31. Forbes-Lorman RM, Harris MA, Chang WS, Dent EW, Nordheim EV, Franzen MA. Physical models have gender-specific effects on student understanding of protein structure–function relationships. Biochem Mol Biol Educ. 2016;44:326–35.
32. Newman DL, Stefkovich M, Clasen C, Franzen MA, Wright LK. Physical models can provide superior learning opportunities beyond the benefits of active engagements. Biochem Mol Biol Educ. 2018;46:435–44.
33. Carlson DL, Marshall PA. Learning the science of research, learning the art of teaching: planning backwards in a college genetics course. Biosci Educ. 2009;13:1–9.
34. Shi J, Wood WB, Martin JM, Guild NA, Vicens Q, Knight JK. A diagnostic assessment for introductory molecular and cell biology. CBE Life Sci Educ. 2010;9:453–61.
35. Couch BA, Wood WB, Knight JK. The molecular biology capstone assessment: a concept assessment for upper-division molecular biology Students. CBE Life Sci Educ. 2015;14:ar10.
36. Newman DL, Snyder CW, Fisk JN, Wright LK. Development of the central dogma concept inventory (CDCI) assessment tool. CBE Life Sci Educ. 2016;15:ar9.
37. Brassil CE, Couch BA. Multiple-true-false questions reveal more thoroughly the complexity of student thinking than multiple-choice questions: a Bayesian item response model comparison. Int J STEM Educ. 2019;6:1–17. https://stemeducationjournal.springeropen.com/articles/10.1186/s40594-019-0169-0
38. Cohen J. A coefficient of agreement for nominal scales. Educ Psychol Meas. 1960;20:37–46.
39. Wright LK, Catavero CM, Newman DL. The DNA triangle and its application to learning meiosis. LSE. 2017;16:ar50.

SUPPORTING INFORMATION
Additional supporting information may be found in the online version of the article at the publisher’s website.

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