

1 Full Title

2 Design and implementation of an asynchronous online course-based undergraduate research experience
3 (CURE) in computational genomics

4 Short Title

5 Asynchronous online research course in computational genomics

6

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33 Abstract

34 As genomics technologies advance, there is a growing demand for computational biologists trained for genomics
35 analysis but instructors face significant hurdles in providing formal training in computer programming, statistics,
36 and genomics to biology students. Fully-online learners represent a significant and growing community that can
37 contribute to meet this need, but they are frequently excluded from valuable research opportunities which mostly
38 do not offer the flexibility they need. To address these opportunity gaps, we developed an asynchronous course-
39 based undergraduate research experience (CURE) for computational genomics specifically for fully-online
40 biology students. We generated custom learning materials and leveraged remotely-accessible computational
41 tools to address two novel research questions over two iterations of the genomics CURE, one testing
42 bioinformatics approaches and one mining cancer genomics data. Here we present how the instructional team
43 distributed analysis needed to address these questions between students over a 7.5-week CURE and provided
44 concurrent training in biology and statistics, computer programming, and professional development. Scores from
45 identical learning assessments administered before and after completion of each CURE showed significant
46 learning gains across biology and coding course objectives. Open-response progress reports were submitted
47 weekly and identified self-reported adaptive coping strategies for challenges encountered throughout the course.
48 Progress reports identified problems that could be resolved through collaboration with instructors and peers via
49 messaging platforms and virtual meetings. We implemented asynchronous communication using the Slack
50 messaging platform and an asynchronous journal club where students discussed relevant publications using the
51 Perusall social annotation platform. The online genomics CURE resulted in unanticipated positive outcomes,
52 including students voluntarily discussing plans to continue research after the course. These outcomes
53 underscore the effectiveness of this genomics CURE for scientific training, recruitment and student-mentor
54 relationships, and student successes. Asynchronous genomics CUREs can contribute to a more skilled, diverse,
55 and inclusive workforce for the advancement of biomedical science.

57 **Author Summary**

58 As technology advances, there is a growing demand for research scientists trained in computational biology but
59 it can be difficult to introduce computer programming and statistics to biology students. One way to meet this
60 demand in an inclusive way is to provide more research opportunities for online students, a significant and
61 growing community which includes many groups underrepresented in the science workforce. We present a
62 course designed for fully-online undergraduate biology students where they can work asynchronously to address
63 a novel research question. We show how we divided research projects among the students of the class,
64 leveraged remotely accessible computational tools and online messaging platforms, and created custom learning
65 materials and assessments to teach the students the necessary biology, computer programming, and
66 communication skills needed for each research project. We demonstrate that students were able to learn the
67 course objectives and cope with academic stresses. Research can be designed around questions in many topics,
68 so we hope that our design can help others to create remote computational research courses in their field.

69

70 Introduction

71 Biomedical science has seen enormous growth in the amount of genomic data produced to investigate the
72 molecular underpinnings of cell biology in health and disease. As high-throughput molecular assays and
73 technology for data processing and machine learning advance, there is an increasing need for cross-disciplinary
74 computational analysts trained to understand biology, genetics, statistics, pharmacology, and mathematical
75 modeling. While typical undergraduate biology courses provide students with a broad background in molecular
76 biology, genetics, and chemistry, many programs still lack substantial instruction in the computation and
77 quantitative analysis (1–3) necessary to analyze genomics data. Barriers to integrating computation include
78 difficulty finding instructors who have had formal training themselves (3,4), so having examples of course
79 materials and techniques for teaching computational genomics and bioinformatics would be beneficial for the
80 genomics research community. In this manuscript, we describe a course format that can be used to successfully
81 teach computational genomics analysis to biology students in a fully asynchronous, online research environment
82 in order to broaden access to research training.

83

84 Course-based undergraduate research experiences (CUREs) are formal courses in which students use well-
85 established scientific practices to participate in novel research projects of interest to the broader scientific
86 community (5). CURE topics are often driven by questions that arise in the instructing faculty's area of interest,
87 thereby providing students with mentors and skills as well as providing mentors with a pipeline to train and recruit
88 students that can advance research programs (6). CURE research projects are aimed at publication, which
89 allows both mentors and students to contribute to the field while allowing the students to gain key research skills
90 and build a science identity (7).

91

92 We designed a course-based research experience (CURE) specifically for fully-online undergraduate students
93 in biology. Online undergraduates are more likely to be from underrepresented demographics in science,
94 including first-generation college students, women, low-income households, and non-traditional adult learners

95 (8,9). Online research experiences have the potential to increase the amount of historically underrepresented
96 student participation in STEM (Science, Technology, Engineering, and Math), increase their application to (and
97 likely enrollment into) graduate programs, and open additional career opportunities, thereby making access to
98 STEM education more equitable and the future workforce more diverse (10–12).

99

100 Bioinformatics and computational biology represent a unique opportunity for the development of fully online
101 CUREs. First, computational analysis resources for -omics (e.g., transcriptomics, genomics, proteomics) level
102 studies are often hosted on high-performance biocomputing clusters, cloud computing environments, or web-
103 based computational analysis platforms whereby users may log in from any location as long as they have a
104 computer and sufficient internet access. Second, transitioning into computational research can be especially
105 challenging for students, perhaps even more so in an asynchronous setting (13,14), but engaging with
106 bioinformatics in a CURE setting can give students more detailed instruction on how to address common sources
107 of anxiety. For Biology students who may not have anticipated learning computational skills, bioinformatics
108 research can include anxiety about a lack of pertinent background knowledge, computer programming anxiety
109 and inexperience, and issues related to accessibility and inclusion in the virtual classroom (13,15). While these
110 factors have been linked to high attrition rates in online STEM courses, the literature shows that retention can
111 increase with student-specific interventions (16), which we aimed to incorporate in this CURE.

112

113 In this manuscript, we describe how we developed and implemented an online CURE in computational genomics
114 to study two different research questions over two iterations of the course. We discuss how the required analysis
115 was distributed among students in an asynchronous format over 7 week-long modules and how research-specific
116 learning materials were designed to promote student success. We demonstrate that our implementation of this
117 CURE led to learning gains among the class and report student responses to research and coding. This
118 presentation of how a computational genomics CURE was designed and implemented for online students is
119 intended to serve as a template for others seeking to expand inclusive and accessible research opportunities at
120 their educational institutions.

121 Course Design and Implementation

122 Ethics Statement

123 Before students began the online genomics CURE, formal written consent was obtained as part of a student
124 experience and demographics survey. This project was conducted with an approved protocol through the
125 Arizona State University Institutional Review Board (approval number STUDY00013025). Students were asked
126 if they were at least 18 years old and if they consent to be part of this study. All students that consented to the
127 study indicated that English was their primary language.

128 Prerequisites and course format for online biology students

129 Students enrolled in this course were online students from a variety of biology majors (Biological/Biomedical
130 Sciences, Neuroscience, Biochemistry, Conservation Biology and Ecology), mostly in the final two years of their
131 undergraduate degree program. Enrollees had interest in computational biology, but most had very little
132 computer programming experience and many had not had opportunities to do research during their online degree
133 program. To give students a basic foundation for computational analysis, students were required to take a
134 prerequisite 7.5-week course in the first half of the semester (BIO 439: Computing for Research; session A in
135 Fall 2022 and 2023). This course assumed no prior coding experience; students were introduced to command
136 line programming, bash scripting, navigating a high-performance computing cluster, and ran through basic
137 genomics analysis (fastq file quality control, alignment, and variant calling). The assignments in the prerequisite
138 course are meant to be a broad first look at how students respond to computer programming and are not specific
139 to the research project chosen for the CURE. All students were given the option of continuing on to the online
140 genomics CURE (session B in Fall 2022 and 2023) but students needed to get approval from the lead instructor
141 before being allowed to continue into the research session.

142 **Research Questions for each CURE**

143 We have implemented two iterations of the online asynchronous genomics CURE covering two separate
144 research questions ([FigureFig 1](#), details in [Supplementary S1 Methods](#)). The pilot iteration of the CURE was
145 based on a bioinformatics project that investigated the effect of sequencing quality on inference of sex differences
146 in gene expression in human placenta. Placenta gene expression data were processed with a range of
147 parameters for sequence quality trimming software and students were asked to compare results to determine if
148 the list of sex differentially expressed genes changed due to the stringency of the parameters chosen. The
149 second iteration of CURE was based on inferring the sex chromosome complement in human cancer cell lines
150 based on expression of genes on the sex chromosomes. Both of these projects were developed organically in
151 the research laboratory of the course instructors, thus ensuring that the instructors had the required expertise to
152 lead the course research and aiding the instructors in communicating why and how the research would contribute
153 to the field.

154 **Distribution of analysis among students**

155 While research training is typically conducted with a one-to-one mentor-student relationship, we employed a
156 paradigm wherein a small team of instructors can provide mentorship to a larger group of students and
157 simultaneously facilitate the development of a network of peers ([FigureFig 1](#)). During the first two modules,
158 course materials and course communications were focused on getting all the students in the class the
159 background information needed to understand the research aims and the way that the research is to be
160 conducted. This included tutorials in the R programming environment, learning how the data were generated,
161 and learning the biological, genomic, and statistical concepts necessary to understand the research aims. In
162 Module 3, all students were assigned the same initial analysis by modifying template code provided by the
163 instructors to facilitate troubleshooting coding errors and streamline guidance on how to interpret the results.
164 For Module 4, students were split into smaller working groups to adapt the code to analyze different but related
165 datasets that address the research question. Instructors assigned student working groups, putting

166 communicative students that were showing technical skills with students that were struggling to maximize the
167 likelihood that all parts of the divided work would be completed successfully and that students had the
168 opportunity to learn from each other. Given small group assignments, students could have someone to work
169 with if they desired or could just as easily work independently; students assigned to the same group could
170 directly compare results and troubleshoot together if needed. Following Module 4, instructors checked data
171 files generated by students for errors and put all the results from all student groups in a shared online location.
172 For Module 5, students were asked to plot and interpret trends across the full data set. This module allowed
173 struggling students time to catch up if they had coding issues in the previous modules and students that were
174 ahead to expand on the analysis they had completed. In Module 6, each student was asked to present their
175 results in the format of a scientific manuscript. Students were allowed to learn from the writing of the other
176 students by conducting a peer review in Module 7. Students were also asked to turn in all code, figures, and
177 output data files so that they can be used to prepare for publication after the completion of the course. In this
178 way, we distributed the analysis, interpretation, and description of the research question equally among the
179 students while simultaneously building in redundancy to make the research goals more robust to individual
180 student challenges.

181

182 Translating Research Plan into Learning Materials

183 Backwards Design of Learning Objectives

184 Learning objectives were written to match the knowledge and skills needed to perform each project within the
185 seven-module format. The main objective for each iteration of the CURE was based on the research project
186 chosen ([FigureFig 1](#)) and then mapped over seven modules (listed in GitHub repository
187 (<https://github.com/SexChrLab/CURES>)). Once the learning objectives were mapped out across the seven
188 modules, module learning pages, reading assignments, and coding assignments were developed to guide the

189 students in achieving those learning objectives. Retention of the most important information and learning of
190 skills in each module was tested using questions in learning assessments (quizzes) during each module.
191 Project descriptions and matching learning objectives for both iterations of the course are available in our
192 **Supplementary-S1 Methods** and GitHub repository (<https://github.com/SexChrLab/CURES>). Instructors
193 analyzed weekly progress reports and assessments to address topics that needed clarification, common
194 misconceptions, and ways to support student learning success in real time (**FigureFig 2**).

195 Module Learning Pages

196 In lieu of expensive textbooks and akin to lab-based research projects, the instruction team wrote collaboratively
197 to produce freely available reference materials to guide the students as they performed the analysis for the
198 chosen research project. Each module had pages posted in the Canvas learning management system. In an
199 introductory module (Module 0), students were introduced to the overall format of the course, introduced to the
200 idea that they will be doing real research where they will collaborate to discover something new, and encouraged
201 to engage in course communications. For Modules 1 to 7, learning pages were written in a modular fashion to
202 keep the format of each module consistent so students could follow them more easily and to increase reusability
203 for future CURE projects. Each learning page had three sections: (1) Biology/Statistics; (2) Coding; and, (3)
204 Professional Development. The Biology/Statistics section was used to describe biological relevance to the
205 analysis, statistics necessary to test hypotheses being generated using the data available, and explain ways to
206 interpret results. The Coding section featured formal instruction on computer programming and explained
207 aspects of the code provided by the instructors. The Professional Development section showed the students
208 ways to seek out information from the literature, described the publication and peer review process, and the
209 cultural norms of scientific research. Additional learning resources for novices and enrichment for more advanced
210 students were included as “Additional Resources”. Published bioinformatics workflows, vignettes, and tutorials
211 were incorporated and/or modified as needed to demonstrate analysis relevant to answering the research

212 question. Module learning pages for both iterations of the course are available in our GitHub repository
213 (<https://github.com/SexChrLab/CURES>).

214
215 Writing learning pages for CUREs took less time with each iteration. For the first iteration, developing the
216 course materials took about one to two weeks per module because the instructors developed all learning
217 materials and assessments completely from scratch while simultaneously deciding on the overall framework of
218 the course. However, developing materials for the second iteration was much quicker (about 3 days per
219 module) because the overall framework for the course was already created and sections of the learning
220 materials could be reused. The Professional Development sections were written to be evergreen and many of
221 the Coding sections could be reused because both iterations of the CURE used code written in the R
222 programming language. As the research being conducted in the CURE is based on the specialty of the
223 instructors, we anticipate that instructors of genomics CUREs will build their own library of learning materials
224 that can be reused, repurposed, or shared with other students for training outside the CURE from year to year.

225 Template Code

226 To facilitate implementing functional code in a short timeline, the instruction team developed template code that
227 the students were asked to modify and build on to complete the analysis required for the research project. Before
228 the course started, instructors processed and prepared the starting input data. For Iteration 1 of the CURE,
229 students were given template code to perform differential gene expression analysis that they could modify to run
230 with different data sets. For Iteration 2, template code was provided to plot expression of a specific gene in
231 cancer cell lines that they could modify to plot other genes and features of interest. Template code was provided
232 in RMarkdown format so that code, products of code, and descriptions of code could be easily created in a single
233 document used for grading, communication, and publication. Template code had many descriptive comments to
234 explain each computational step and model best practices for coding technique. Tutorials assigned at the
235 beginning of the CURE were chosen based on how well they explained the features in the template code so that

236 students would feel more confident in making changes to suit the purposes of the research. Template code for
237 each iteration of the course is provided in Github (<https://github.com/SexChrLab/CURES>).

238 **Student assessments and surveys**

239 **Pre- and post- learning assessments**

240 Students took an identical learning assessment before and after completing the genomics CURE ([FigureFig 2](#)).
241 Questions were designed by the instructors to match the learning objectives of the course in Biology/Statistics,
242 Coding, and Professional Development, and vetted by outside genomics experts. Likert surveys were included
243 to gauge student comfort levels with specific skills taught in the genomics CURE. Scores and answers were not
244 revealed to the students and students received full credit regardless of the percentage of correct answers.

245 **Weekly progress reports**

246 To maintain an authentic research experience, the instruction team developed an open-response summative
247 assessment for students to submit weekly, designed in the fashion of progress reports submitted in the lead
248 faculty instructor's laboratory ([FigureFig 2](#)). To focus the students' attention on research progress (not just
249 grades on assessments), the progress report prompted students to list their accomplishments, challenges they
250 faced, and how they addressed those challenges. The second iteration of the CURE also included a scientific
251 writing prompt, wherein students were asked to describe specific methods and results each week, receiving
252 iterative feedback prior to compiling their final manuscript report.

253 **Coding assignments**

254 Research milestones were assessed using weekly coding assignments, typically requiring upload of code reports
255 and data output files. Feedback was provided to students through the Canvas learning environment using a
256 custom rubric highlighting key concepts. Because more value was placed on effort and progress than on specific

257 outcomes, the progress report was worth 100 out of 125 points of the weekly module assignment grade and
258 coding assignments were only worth 25 points.

259 **Manuscript and peer review**

260 As a final project in the CURE, students were asked to write a report in the format of a manuscript for publication
261 and then asked to conduct peer reviews on manuscripts written by their classmates ([FigureFig 2](#)). This trained
262 students to communicate scientific results with the level of accuracy and detail expected at a professional level.
263 The professional development section of several modules was used to walk students through how to put together
264 the various parts of their manuscript. Students were taught how to write detailed legends for figures produced
265 as the research project was conducted in Modules 3, 4, and 5, then asked to create a storyboard outline of these
266 figures to weave a complete set of related analyses that address the research project aims in Module 5, before
267 fully writing about these results in Module 7. Examples of how to keep track of and document software packages
268 used in code in the methods section of the manuscript were featured in all template code. Class discussions on
269 topics involved in the research helped develop context that was featured in the abstract, introduction, and
270 discussion portions of the manuscript. Many students wrote about results that were unexpected as those results
271 fueled many discussions in lab meetings and Slack throughout the course, such as having a lot of overlap in
272 results with different trimming parameters in Iteration 1 and cancer cell lines having sex chromosome gene
273 expression that did not match what was expected based on the reported sex of the patient from which the cancer
274 cell line was derived in Iteration 2. As all students did similar analyses, they were able to appreciate the
275 descriptions and insights offered by their classmates during the peer review process. The rubric used to grade
276 peer reviews of manuscripts is in the provided Github repository (<https://github.com/SexChrLab/CURES>).

277 **Class Communication**

278 The primary mode of communication with students was the Slack messaging system so that responses to
279 questions and posts were visible to all students. This platform was integrated into the Arizona State University

280 Canvas learning management system and allowed for asynchronous communication between students (see
281 [**Supplementary-S1 Methods**](#) for details and alternatives). Students were encouraged to post and respond to
282 Slack messages about confusing concepts, coding problems, and any other challenges encountered during the
283 course. Instructors posted about common misconceptions, bug fixes, activities and resources for learning
284 enrichment, and encouragement for students as they struggle with challenging material. In addition to Slack,
285 students were given several opportunities each week to meet with the instruction team synchronously: lab
286 meeting, writing hours, and shared research hours ([**FigureFig 2**](#)). Optional weekly lab meetings recorded and
287 transcribed in Zoom covered student-reported challenge topics. Optional weekly writing hours gave students the
288 opportunity to discuss interpretation of findings they would write about in their weekly assignments and
289 manuscript. Optional shared research hours were offered twice a week, giving students the opportunity to
290 troubleshoot code with the instruction team live with options to share their screen.

291 Asynchronous Journal Club

292 In the second iteration of the CURE, instructors implemented an asynchronous journal club to help students
293 engage with peer-reviewed literature relevant to the research project ([**FigureFig 2**](#)). To do this, the instructors
294 chose one publication each week for the first five modules for the students to read (leaving the last two modules
295 with more time to focus on the manuscript and peer review). To encourage collaborative learning, the chosen
296 publications were posted using the Perusall social annotation environment and students were given points
297 toward their final grade for posting comments as they read the paper (see [**Supplementary-S1 Methods**](#) for more
298 information). We found that the students pointed out what they thought was interesting or relevant about the
299 publications and asked and answered each other's questions about the journal articles. This format of journal
300 club provided the flexibility needed for online student learning while providing asynchronous but meaningful
301 discussions about important published works in the field.

302 Grading and instructor communications

303 Custom rubrics and examples were provided to students to describe expectations for assignments and
304 scientific writing. The majority of the course grade for the CURE was based on the weekly progress report so
305 the majority of the grading efforts of the instructors were dedicated to this ([Figure Fig 2](#)). As the progress report
306 was divided into sections, instructors were able to split the work, typically with one instructor grading the
307 accomplishments and challenges sections and another grading the scientific writing and coding assignments
308 addressing the research aims. Instructors held a short weekly meeting to discuss research progress and how
309 to address student challenges. Since a great deal of the grade for the CURE was based on open-response
310 writing, instructors watched carefully for signs of academic dishonesty, including inappropriate use of artificial
311 intelligence chatbot engines and copying between students ([Supplementary S1 Methods](#)), but because so
312 much emphasis was placed on progress over products, the vast majority of students submitted their own work
313 and experiences throughout the CURE.

314 Assessing student learning and student comfort

315 A significant improvement in overall scores between pre- and post-assessments was observed in both iterations
316 of the genomics CURE. The first iteration of the CURE showed a significant increase of 11.64% in the mean
317 score from 65.96% to 77.61% (n = 13 students, p = 0.003, [Figure Fig 3A](#)). This is considered to be a medium to
318 large effect size as evaluated by Cohen's d-statistic (d = 0.79, medium effect size range = 0.5 - 0.8, large effect
319 size range > 0.8). The second iteration of the CURE showed an increase of 3.5% in the mean score from 61.62%
320 to 65.16% but this increase was more statistically significant and deemed a large effect size given a much larger
321 class size (n = 45 students, p = 8x10⁻⁷, d = 0.82, [Figure Fig 3B](#)). Students consistently showed the most
322 significant increase in the Biology/Statistics section in both iterations of the CURE ([Figure Fig 3C, and](#)
323 [Supplementary Figure 2 Fig](#)), while the Coding and Professional Development sections showed small to
324 medium levels of effect on student learning. The learning assessment questions for the second iteration were

325 also given a subtopic within Biology/Statistics, Coding, and Professional Development, which shows that the
326 increased learning in Professional Development in the second iteration ([Figure Fig 3C](#)) was driven by a
327 significant improvement in the subsection about reading scientific papers ([Supplementary Figure S3 Fig](#)). This
328 is likely due to the implementation of the asynchronous journal club specifically in Iteration 2 of the online
329 genomics CURE. Progress reports for students whose learning assessment scores did not improve after taking
330 the CURE included reports of unexpected issues in their personal lives as well as some who were unable to
331 make the necessary learning gains required to understand and implement the code. Likert questions included
332 on the learning assessment also showed increased comfort with the skills taught during the CURE for many
333 students ([Supplementary Figure S4 Fig](#)).

334 Understanding student experiences

335 To more thoroughly understand student experience throughout the CURE and increase student engagement,
336 each week the instruction team reviewed the progress report for challenges reported and how the student
337 responded to or coped with those challenges. Summaries of the progress reports ([Figure 4–A-C and](#)
338 [Supplementary Figure S1](#)) show that students frequently felt challenged by needing to understand the material,
339 writing and troubleshooting code, and managing their time to complete the research aims while having a full
340 course load. Time was mentioned mostly frequently in the exploration and introduction phase (Modules 1 and 2)
341 as it is challenging to jump into a research project; many students wrote about searching for more information to
342 fuel their curiosity beyond the learning materials. In the analysis phase of the research (Modules 3, 4, and 5),
343 coding became the most frequent reported challenge. Reading details about what aspect of the coding was the
344 most challenging for students allowed the instructors to provide support to students to help solve common issues
345 and to improve coding tutorials in future iterations of the CURE. Additionally, many students reported in their
346 progress reports that they had full-time jobs and care-taking responsibilities that contributed to difficulty with time
347 management. In response to time management concerns, instructors provided learning materials in smaller
348 sections and provided time estimates where possible to help students manage their work time effectively. In the

349 analysis modules, students commonly mentioned Slack, and asked for help understanding errors while compiling
350 the R Markdown report, demonstrating that they looked to collaborate to solve coding problems. In the reporting
351 or manuscript phase (Modules 6 and 7), students reported challenges with writing and getting information and
352 figures into their manuscript. Many students reported that this was the first time they were asked to synthesize
353 new results instead of applying techniques indicated by instructors to get a predetermined result. Some weren't
354 sure if they had done enough to address the research aims or if their writing was clear enough; instructors were
355 able to provide feedback and resources to address these concerns. Many students reflected on the quality of
356 their own work after reading other students' work in the peer review in Module 7. Analysis of general trends on
357 how students coped with challenges showed that a high proportion of students employed adaptive coping
358 strategies (problem solving, support seeking, information seeking, and self-reliance) throughout the course as
359 these were encouraged and rewarded, while the rate of maladaptive coping strategies reported increased
360 towards the end of the course as students worked hard to describe results in their final manuscript and had
361 struggled to keep up with final assessments while also managing the rest of their academic schedules
362 ([Supplementary Figure S5 Fig](#)). Select statements from students' progress reports can be found in the
363 **Supplemental S1 Methods**. These statements demonstrate that while students struggled trying to learn skills
364 and knowledge they felt were interesting and important, ultimately, they were able to achieve meaningful growth
365 in their research mindset and overall knowledge of the topics being studied. Progress reports provided a way for
366 instructors to look beyond students' grades to follow their journey through the research project while learning to
367 manage the uncertainty that comes with doing research. Progress report helped instructors provide feedback
368 and resources tailored to the needs of the students in each iteration of the CURE.

369 Discussion and Future Directions

370 Upon completion of the genomics CURE, all students were invited to continue to work on the research project
371 the following term. Many students requested to continue on with the project by doing follow-up analysis or present
372 the genomics results for research symposiums and publication. Follow-up studies focused on assessing the

373 generalizability of the results from the CURE with other related datasets and analytical tools. In progress reports
374 and course discussions, students discussed seeking out research and professional development opportunities
375 outside of the CURE research project. Additionally, mentors may be able to recruit students as more permanent
376 members of their laboratory. This study demonstrates that students can be successful in online research
377 experiences that incorporate accessible learning materials, multiple options for class communication, and open-
378 response progress reports to monitor achievements, challenges, and coping strategies.

379

380 By directly assessing student learning and experiences throughout the CURE, the instructors were able to tell
381 which areas of the course are effective and which to prioritize for improvement. Based on the feedback from
382 students, instructors continually improved the introductory materials for the earlier modules in the course to help
383 ease students into the research project and coding. Instructors are currently developing better ways to share
384 communication and discussion between students including a bulletin board Slack channel where instructors can
385 post results and accomplishments for all students to learn from throughout the course and provide more detailed
386 templates for the final manuscript and other assignments to communicate expectations more clearly. Instructors
387 are investigating more automated ways to collect and analyze information from the progress reports to make
388 grading more scalable and inform discussion topics and interventions. An instructor handbook is being created
389 to help share lessons learned over many iterations of the CURE.

390

391 Opening genomics research opportunities to online students using an asynchronous format allows many
392 students who would typically be excluded to participate in important research endeavors. Asynchronous
393 genomics CUREs are a way to bring valuable research opportunities, mentorship, and analytical skills to many
394 students who can go on to contribute to a more diverse and capable workforce to tackle an ever-expanding set
395 of genomics challenges.

396

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448

449 Figures

450 Figure Legends

451 **FigureFig 1: Project summaries and distribution of analysis among students.** Two iterations of the online
452 genomics CURE were taught in the same format and used to conduct two completely different research
453 projects. The first two modules were dedicated to exploration of the data set being analyzed and featured an
454 introduction to the coding language being used in the course. The middle modules were used to divide the
455 required analysis between the students of the class. For iteration 1, students studied how parameters for
456 quality trimming of RNA sequencing data affected identification of genes differentially expressed by sex in the
457 human placenta. For iteration 2, students used expression of specific sex chromosome genes to infer the sex
458 chromosome complement in cell lines used as models for human cancer. In the last modules, each student
459 described the results of the study in a manuscript which was peer reviewed by other students.

460 **FigureFig 2: Instructor and student contributions over 7-week CURE.** Instructor contributions are labeled
461 as circles, student contributions in triangles. Students began by filling out a pre-assessment to show their
462 baseline level of knowledge on the research topic. Each week, students completed research goals and turned
463 them in as assignments along with a progress report communicating their achievements and challenges.
464 Students were assigned weekly scientific writing prompts and publications to read in a journal club designed to
465 help them slowly build up their final reports in Module 6. Instructors used all of these submissions to select
466 topics for weekly lab meetings conducted using teleconference software and recorded for students that could
467 not attend. Instructors hosted shared research (office) hours throughout the week. Instructors monitored
468 progress on research goals and distributed data generated by individual students at the midpoint of the CURE.
469 Following the completion of the course, students completed a learning assessment which instructors analyzed
470 to see which areas students were able to increase their knowledge in as well as misconceptions and struggles
471 that could be avoided in future CUREs.

472 **FigureFig 3. Increase in student knowledge and skills after genomics CURE.** (A,B) Boxplots depicting mean
473 student assessment scores before (green) and after (orange) completing the genomics CURE. Each point
474 represents a student who completed both the pre- and post- assessments and the lines connect pre-assessment
475 and post-assessment scores for each student. The mean class score significantly increased from 65.96% to
476 77.61% (paired t-test p value 0.003, Cohen's D-statistic 0.79, medium-large effect) for iteration 1 (A) and 61.62%
477 to 65.16% (paired t-test p value 0.00002, Cohen's D-statistic 0.82, large effect) for iteration 2 (B). (C) Boxplots
478 depicting each pre-assessment (green) and post-assessment (orange) score for all questions divided by topic
479 for CURE Iteration 2: Biology/Statistics, Coding, and Professional Development.

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FigureFig 4. Challenges across CURE modules (Iteration 2). (A-C) Word cloud summary of challenges reported in
weekly progress reports in three main research phases: Module 1 (A) representing the data exploration and coding
introduction phase, Module 3 (B) representing the analysis and interpretation phase, and Module 6 (C) representing the
reporting results phase. Word use frequency is shown by size and color (larger and darker shade of red for higher
frequency).

489 Supporting Information

490 491 Supplementary Figures

492
493 **Supplementary S1 FigureFig 1: Summary of accomplishments and challenges section of the weekly**
494 **progress reports for Iteration 2 of the CURE.** For each module, word clouds are provided to summarize
495 what students were accomplishing given the research aims (provided on the left in blue) and what challenges
496 were encountered. Word size and color is used to highlight high frequency words, larger and dark shade of red
497 indicating high frequency.

498
499 **Supplementary S2 FigureFig 2: Increase in student knowledge and skills after genomics CURE by**
500 **Topic (Iteration 1).** Boxplots depicting each pre-assessment (green) and post-assessment (orange) scores for
501 all questions divided by topic for CURE Iteration 1: Biology/Statistics, Coding, and Professional Development.

502
503 **Supplementary S3 FigureFig 3: Increase in student knowledge and skills after genomics CURE by**
504 **Subtopic (Iteration 2).** Violin plots depicting each pre-assessment (green) and post-assessment (orange)
505 scores for all questions divided by subtopic for CURE Iteration 2. Subtopics showing paired t-test p-value less
506 than 0.01 are highlighted in red, between 0.01 and 0.05 in orange, and between 0.05 and 0.1 in yellow.

507
508 **Supplementary S4 FigureFig 4: Questions about student comfort level in specific skills in the**
509 **genomics CURE from pre- and post- learning assessments.** Students were asked to rate their comfort
510 level on a scale from very uncomfortable to very comfortable (6 options total, responses assigned numerical
511 values between -3 and +3) for skills used throughout the course: programming in R, reading and writing
512 scientific papers, asking questions about coding in a class setting, and using command line programming in a
513 Linux environment. Boxplots depicting each pre-assessment (green) and post-assessment (orange) scores for
514 all 5 questions for all students are shown with a paired t-test p-value showing the statistical significance of the
515 improvement after the completion of the CURE.

516
517 **Supplementary S5 FigureFig 5: Trends in coping strategies across CURE modules.** Proportion of students
518 reporting various coping strategies to overcome challenges encountered during genomics research. Responses for
519 progress reports for each module were categorized as adaptive, maladaptive, or those that could be either depending on
520 context. Adaptive themes include problem solving (red), support seeking (orange), information seeking (gold), self-
521 reliance/emotional regulation (olive), and cognitive restructuring (green). Maladaptive themes include escape (light blue),
522 isolation (blue), rumination (purple), helplessness (lilac), delegation (fuchsia), and opposition (pink).

523 Supplementary S1 Methods

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525
526 This section provides additional details on the research projects chosen for both iterations of the genomics
CURE, instructions given to instructors and students to promote a collaborative research environment,
implementation of various tools used for instruction, and insights that could be helpful when creating a CURE.