(1) Major activities.

The budget period for this report was 09/21/2017 – 06/30/2018. In this budget year, major activities of the project primarily focused on administrative activities to mobilize the collaborating teams (4 national, 1 international). In following, the teams involved in efforts to establish consensus on the detailed workflow of the project and to start modeling and simulation activities along with community outreach.

(2) Specific objectives.

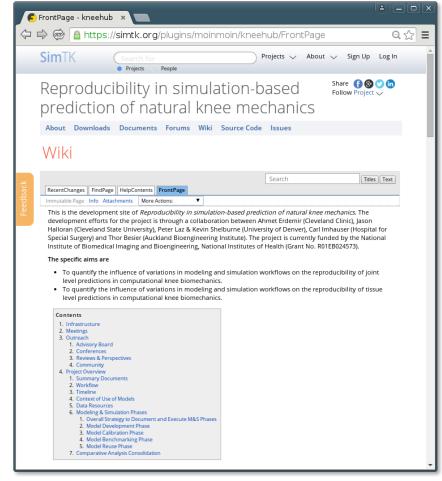
The overall goal of the project is to understand how modelers' choices to build models, even when using the same data, may influence predictions and therefore the reproducibility of simulation results. Specific aims are i) to quantify the influence of variations in modeling and simulation workflows on the reproducibility of joint level predictions in computational knee biomechanics, and ii) to quantify the influence of variations in modeling and simulation workflows on the reproducibility of tissue level predictions in computational knee biomechanics.

The primary aim of the reporting period was the organization of five collaborating teams – Cleveland Clinic, Cleveland State University, University of Denver, Auckland Bioengineering Institute, Hospital for Special Surgery; administratively, contractually, and from a scientific workforce perspective. Specific to the modeling and simulation activities, the goals were to decide on the specific data sets to utilize from existing data resources and to start model development activities.

(3) Significant results, including major findings, developments, or conclusions.

Administratively, all subcontracts were setup, albeit in a delayed manner, and teams have started recruiting research fellows, graduate students, and engineers. Online infrastructure for collaboration and outreach has been maintained [1]. Collaborating teams had a site visit at the Cleveland Clinic to start the project and have multiple meetings, some web-based and one during the annual meeting of the Orthopaedic Research Society

(ORS). The exact nature of modeling and simulation phases and the timeline were decided: model development. calibration, model benchmarking, and model reuse. Specimens from data resources were also agreed upon: oks003 from Open Knee(s) and DU02 from Natural Knee Data. Instructions on the development phase disseminated to each team and the public on the project wiki [2]. Currently, the teams are at the model development phase. Data earmarked for the model phase development were disseminated publicly at the project site the teams to utilize [1]. investigators also reached out to the thirdparty reviewer, Tina Morrison from US Food and Drug Administration, to explain project deliverables for credibility and reproducibility assessment and the teams' from comparative expectations the analysis. Community outreach has been evolving organically; the investigators delivered a group presentation at the Clinic on "art" Cleveland the of computational modeling in knee biomechanics. At the ORS 2018 annual meeting, team leaders engaged with



scientists of knee biomechanics and of **Figure 1.** Project wiki provides the gateway for collaborating teams and reproducibility in science. The teams are the public access to information on project phases [2].

tentatively planning to have a scientific workshop at ORS 2019. They are also in the process of writing a manuscript on modeling and simulation workflows in knee biomechanics.

(4) Key outcomes or other achievements.

The project website is fully operations [1]. The wiki provides a gateway for the collaborating teams and the public to get instructions on project phases and links to data resources and in future, modeling and simulation specifications and outputs [2] (Figure 1). Dissemination of data earmarked for modeling phases has already started in the "Downloads" section of the project site [1].

The modeling teams, along with the third-party reviewer, presented a study on comparison of the project workflow to broadly applicable guidance on credible practice of modeling and simulation [3]. To promote the project in the European community, an abstract was submitted to VPH 2018, Virtual Physiological Human Conference to be held in Zaragoza in September 2018. The Principal Investigator has also been involved in synergistic scholarly activities; on opinions on model sharing and reproducibility in the biomechanics community [4] and on credibility of simulation in neurosciences [5].

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