

METHOD ARTICLE

Organizing and running bioinformatics hackathons within Africa: The H3ABioNet cloud computing experience [version 2; peer review: 2 approved, 1 approved with reservations]

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Abstract

The need for portable and reproducible genomics analysis pipelines is growing globally as well as in Africa, especially with the growth of collaborative projects like the Human Health and Heredity in Africa Consortium (H3Africa). The Pan-African H3Africa Bioinformatics Network (H3ABioNet) recognized the need for portable, reproducible pipelines adapted to heterogeneous computing environments, and for the nurturing of technical expertise in workflow languages and containerization technologies. Building on the network's Standard Operating Procedures

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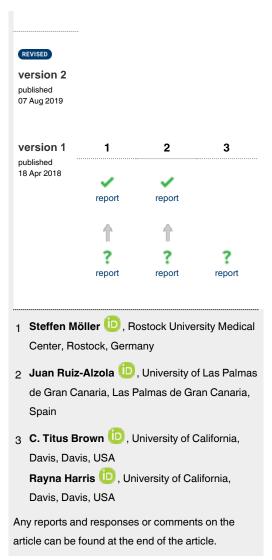
(SOPs) for common genomic analyses, H3ABioNet arranged its first Cloud Computing and Reproducible Workflows Hackathon in 2016, with the purpose of translating those SOPs into analysis pipelines able to run on heterogeneous computing environments and meeting the needs of H3Africa research projects. This paper describes the preparations for this hackathon and reflects upon the lessons learned about its impact on building the technical and scientific expertise of African researchers. The workflows developed were made publicly available in GitHub repositories and deposited as container images on Quay.io.

Keywords

Bioinformatics, hackathon, workflow, reproducible, pipeline, capacity building



This article is included in the African Society of Human Genetics gateway.



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REVISED Amendments from Version 1

We would like to extend our sincerest gratitude to the reviewers for their comments and constructive criticism on our article entitled "Organizing and running bioinformatics hackathons within Africa: The H3ABioNet cloud computing experience". We have carefully and thoroughly evaluated all the comments and addressed them as necessary in the current version of our revised article. We do hope that we have tackled the issues raised in the comments to standards that meet your approval. Below is a brief summary of the main revisions to our article:

We have changed the URL for the SOPs site for common data analysis tasks within the H3ABioNet consortium from http:// h3abionet.org/tools-and-resources/sops to https://h3abionet. github.io/H3ABionet-SOPs/index.html. We have added a new section "Context, rationale and impact" to provide a better explanation of the context of the hackathon, H3ABioNet and H3Africa in Africa. The "Post-hackathon feedback and actions" section title has been revised to "Post-hackathon activities". A paragraph has been added to the "Discussion" section, which discusses the context and factors enabling the success of the hackathon in light of limitations to infrastructure and access to internet resources in Africa. The legend on Figure 1 has been updated with a better description of the key points. A new table has been added to the article (Table 1) to highlight the significance of the pipelines. The table summarizing the different communication channels (Table 1 in the original article) has been updated and is now Table 2. Six new references used in the revision of the article have been added. A few syntax changes were in order for better clarity.

We would like to once again express our gratitude and appreciation to the reviewers for their comments on our article. Please feel free to contact us for any further queries.

See referee reports

Introduction

As an inherently interdisciplinary science, bioinformatics depends upon complementary expertise from biomedical scientists, statisticians and computer scientists¹. This opportunity for collaborative projects also creates a need for avenues to exchange knowledge¹. Hackathons, along with codefests and sprints, are emerging as an efficient means for driving successful projects². They can be in the form of science hackathons that aim to derive research plans and scientific write up³, community-driven software development⁴, and data hackathons or datathons⁵. In addition to the scientific and technical outcomes, these intensive and focused activities offer necessary skills development and networking opportunities to young and early career scientists.

On the African continent, there is generally limited access to such events. However, with the growing capacity for Africans to generate genomic data, the need to analyze these data locally by African scientists, is also growing. H3ABioNet⁶, the Bioinformatics Network within the H3Africa initiative⁷, has invested in capacity building via different approaches⁸. The H3ABioNet Cloud Computing hackathon was a natural extension of the network's efforts in developing Standard Operating Procedures (SOPs) via its Network Accreditation Task Force (NATF)⁶; aimed at building and assessing capacity in genomic analysis. This also follows other efforts by the H3ABioNet Infrastructure Working Group (ISWG) towards setting up infrastructure at various H3ABioNet Nodes at the hardware.

software, networking, and staff level. The H3ABioNet Cloud Computing hackathon, therefore, provided an excellent opportunity to assess the computational skills capacity development of the network through training, learning and adoption of novel technologies (Figure 1). These technologies included workflow languages for reproducible science, containerization of software, and creation of computational products that can be used in heterogeneous computing environments encountered by African and international scientists in the form of standalone servers, cloud allocations and High-Performance Computing (HPC) resources.

In this paper, we discuss the organization of the H3ABioNet Cloud Computing hackathon, the interactions between the participants, and the lessons learnt. Baichoo *et al.*¹⁰ describe the technical aspects of the pipelines, whereas the code and pipelines themselves have been made publicly available via H3ABioNet's GitHub page in the following repositories: (h3agatk, h3abionet16S, h3agwas and chipimputation) as well as container images hosted on Quay.io.

Context, rationale and impact

For a healthy and strong scientific community, knowledge sharing activities, such as hackathons, are paramount. While instrumental to collaboration and efficient in developing solutions to shared problems, such activities are limited within Africa.

The H3ABioNet consortium aims to build a coherent and strong bioinformatics community within Africa that can technically support H3Africa projects for within-Africa analysis of African data. A network of > 27 nodes, H3ABioNet unites researchers from 15 African countries, in addition to a node in the US. Establishing a baseline where each node had sufficient computational infrastructure to carry genomics analyses was (and still) one of the key deliverables of the consortium. Consortium projects like Netmap helped to achieve this goal by evaluating network connectivity between the participating nodes and also led to upgrading infrastructure where warranted¹¹.

Consequently, the primary value of the H3ABioNet cloud computing hackathon was to expose African scientists to the practical aspects of community development of computer code and to try to create a community around the maintenance of a set of workflows that implement methods that are useful to the H3Africa research community and beyond.

More pragmatically, the workflows developed in the hackathon serve as practical implementations of the Standard Operating Procedures for the H3ABioNet Accreditation Exercises, which are used to evaluate the capacity of African research groups in analyzing complex genomic datasets- like those being produced by various H3Africa research projects⁹. Success in taking one of the exercises is considered a landmark for African groups who are preparing to step into the existing gap between data production and data analysis, where the analysis is typically undertaken by First World groups.

Today, those implemented pipelines have been used for data analysis within the context of H3Africa projects, and/or

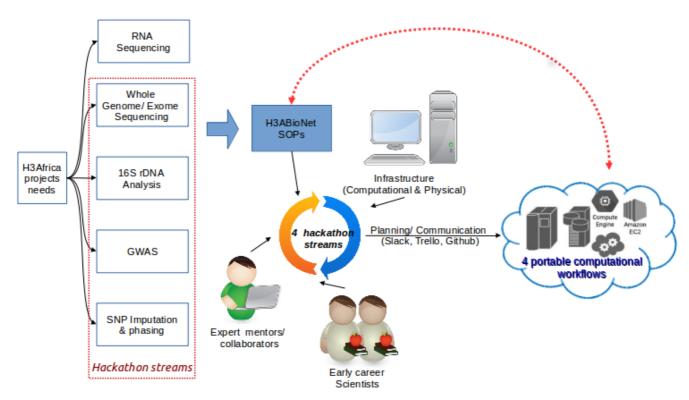


Figure 1. Planning and execution of the H3ABioNet Cloud computing hackathon. H3ABioNet developed SOPs for 5 analysis niches needed within H3Africa projects. 4 out of these were implemented as portable workflows as a result of the H3ABioNet 2016 Cloud Computing hackathon that brought together early career scientists, expert mentors and collaborators by utilizing many planning and communication platforms. (SOPs: Standard Operating Procedures).

incorporated into H3ABioNet training materials. Table 1 below highlights the significance of each developed pipeline, along with some technical notes about its implementation and availability. An extensive technical evaluation and trajectory of development is found in 10.

H3ABioNet Cloud Computing Hackathon Activities

Prior to the H3ABioNet Cloud Computing hackathon, H3ABioNet, via its Infrastructure Working Group (ISWG), formed a Cloud Computing task force to investigate cloud computing technologies, familiarize H3ABioNet members with current cloud implementations and gauge their suitability for H3Africa data analyses. The H3ABioNet Cloud Computing hackathon was one of the first deliverables of this task force, with the specific objective to test and implement four analysis workflows that can be ported on multiple computing platforms. Figure 1 shows this hackathon within the broader H3Africa context and provides a broad overview of the planning and execution of this activity, with details in the following subsections.

Pre-hackathon preparations

The computational pipelines put forward for development during the H3ABioNet Cloud Computing hackathon were identified based on the data being generated by different H3Africa projects and the SOPs used for the H3ABioNet Node

Accreditation exercises. Reproducibility and portability were also identified as key features for the workflows, due to the heterogeneous computational platforms available in Africa. H3ABioNet Nodes that used or helped develop current H3ABioNet workflows and SOPs were part of the planning team, as well as other nodes that had technically strong scientists who were willing to extend their skills.

In the course of planning for the H3ABioNet Cloud Computing hackathon, two technical areas were identified where additional expertise was required. These were containerization technology such as Docker, and the writing of genomic pipelines in popularly used workflow languages and newly emerging community-standards like Nextflow¹² and the Common Workflow Language (CWL)¹³, respectively. While expertise for Nextflow already existed within the network, two collaborators from outside Africa were interested to join the project given their expertise in cloud environments, containerization of code¹⁴ and developing CWL¹³. They subsequently joined the planning and participated in the hackathon. In fact, they were also invited as guest speakers in the network's monthly webinar series where they shared some of their experiences in these areas with the broader H3ABioNet consortium.

The H3ABioNet Cloud Computing hackathon was announced on the internal H3ABioNet consortium mailing list as a call

Table 1. Significance and impact of the developed pipelines as part of the H3ABioNet 2016 Cloud Computing hackathon, along with implementation notes.

GitHub link**	https://github.com/h3abionet/h3agatk	https://github.com/h3abionet/h3abionet16S	http://github.com/h3abionet/h3agwas	https://github.com/h3abionet/chipimputation/
Testing environment*	 EGI FedCloud resource (+) AWS ec2 (+/-) Microsoft Azure VM (+/-) 	 AWS EC2 & Azure VMs (+/-) SGE cluster (+) PBS cluster (-) 	PBS cluster the Bright Cluster Manager (-) AWS EC2 (Docker Swarm and cloud-init)	SGE cluster (-) OpenStack cloud (+)
Implementation Significance & Impact	Such data is extensively generated within H3Africa projects (for example, the data informing the design of the African Genotyping chip ¹⁵ enriched by variants from 350 deeply sequenced African genomes)	For performing 16S rDNA diversity analysis of microbial species in metagenomic samples (was derived from work done to analyze bacterial populations present in leg ulcers of sickle cell patients in Nigeria.)	The H3Africa Consortium will genotype over 30,000 individuals using a custom designed African genotyping array. H3Africa projects, like AWI-Gen¹6 have already extensively used this pipeline to analyze more than 11.5k samples at the time of writing. Additionally, this pipeline is now part of H3ABioNet training resources on GWAS, with online content readily available via (https://www.youtube.com/playlist?list=PLcQ0XMykNhCQJPz0a mnbz9BPM4Bu0Nkgf); and also for in-person, "Bring your Own Data" workshops	Of value in population structure and admixture studies. Eventually, this pipeline (along with computational resources from well-resourced H3ABioNet nodes) are intended to be provided as a service to African researchers. Currently, this pipeline too is part of H3ABioNet training resources on GWAS, with online content readily available via (https://www.youtube.com/playlist?list=PLcQ0XMykNhCQJPz0amnbz9BPM4Bu0Nkgf); and also for in-person, "Bring your Own Data" workshops
Implementation	CWL	CWL and Nextflow	Nextflow	Nexflow
Analysis pipeline	Whole Genome/ CWL Exome NGS Data Analysis	16S rDNA Diversity Analysis	Genome-wide association studies (GWAS)	SNP imputation Nexflow

^{* +} and - indicates testing with and without docker, respectively, in the given environment

^{**} Corresponding docker containers are available at: https://quay.io/organization/h3abionet_org and https://dockstore.org/workflows/h3abionet/h3agatk

for interested applicants and in some cases, individuals were invited based on their specific expertise. Most of the participants selected were early career scientists with strong computational skills, an understanding of genomic pipelines and willingness to work in teams. The pipelines for the Cloud Hackathon were divided into four "streams": 1) Stream A: variant calling from whole genome sequencing (WGS) and whole exome sequencing (WES) data (https://github.com/h3abionet/h3agatk), 2) Stream B: 16S rDNA Diversity Analysis (https://github.com/ h3abionet/h3abionet16S), 3) Stream C: Genome Wide-association studies (Illumina array data) (https://github.com/h3abionet/ h3agwas) and 4) Stream D: SNP Imputation and phasing using different reference panels (https://github.com/h3abionet/ chipimputation). Successful applicants were given a choice to select a project stream based on their skills and interest- or if unsure, assigned to a specific stream. Streams A and B decided to use CWL for their pipeline development, whereas Streams C and D opted to use Nextflow due to their prior experience using Nextflow.

Stream membership respected participants' own interests, but it was also sought to have steams of balanced composition. This included bioinformaticians with knowledge in the specific genomic analyses and computational tools required, strong computational skills to create the Docker containers and implement workflows, and strong system administration skills to assist with the installation of numerous software components as needed. We also included bioinformaticians with experience in running the workflows or components of the workflows, and software developers who could assist with creating Docker containers, troubleshoot and implement workflow languages (CWL was still in draft-2 at the time of the hackathon, and some language features were added based on our experience).

To maximize the learning experience, upon selection, participants were given prerequisite tutorials and materials (Github, Nextflow, CWL, Docker and the SOPs) to go through. Communication and planning infrastructure in the form of Slack channels and Trello boards were created beforehand with all the participants added in order to allow them to brainstorm and share ideas with team members before the hackathon began (Table 2). Fortnightly planning meetings were held starting from 3 months in advance in order for hackathon participants to

get involved in planning their proposed tools and to get to know one another and develop a working rapport before the start of the hackathon.

The hackathon ran in August 2016 and was hosted at the University of Pretoria Bioinformatics and Computational Biology Unit in South Africa. The choice of the hackathon venue was based on the availability of Unix/Linux desktop machines with the facility for sudo/root access enabling participants to install software and deploy Docker containers for testing. Besides the local machines, participants also had access to cloud computing platforms such as Azure and Amazon, Nebula (made available by the National Center for Supercomputing Applications, University of Illinois at Urbana-Champaign), and the African Research Cloud (through a collaboration with the University of Cape Town eResearch initiative). After the hackathon, more testing was also done on EGI Federated Cloud resources (as a courtesy allocation to the University of Khartoum).

Hackathon week activities

The initial day of the H3ABioNet Cloud Computing hackathon was dedicated to introductions, expectations by the participants and practical tutorials covering the use of CWL, Nextflow and creation of Docker containers to ensure all participants had the same basic level of knowledge. The teams had a breakout session where overall milestones for the streams during the hackathon week were refined, tasks were identified and assigned to team members and Trello boards updated with the specific tasks. Each stream reported back on their progress and overall work plan for the coming hackathon days. For the remaining days of the hackathon, participants were split into their respective streams to work on developing and containerizing their pipelines as well as creating the related documentation. To ensure a successful hackathon with concrete outcomes, the streams spent the first 30 minutes of each hackathon day reviewing their prior progress and updating their Trello boards and reporting to the group what they will be working on. At the end of the day, each stream provided a progress report to the whole group on what they had achieved, what they struggled with and what they will be working on. The start and end of day reporting proved useful as it allowed groups that had encountered and solved an issue to share the implemented solution with another stream, and for different streams to work together

Table 2. Communication channels used for the hackathon.

Channel	Link	Purpose
Mailing list	-	Group wide announcements and communications
Moonf	https://mconf.sanren.ac.za/	Online meetings
Slack	https://slack.com/	Inner group discussions and chat
Trello	https://trello.com/	Plan goals and activities, and track progress
GitHub	https://github.com/	Code repository and version control
Google Drive	https://drive.google.com/drive	Document sharing

to solve any shared issues encountered, thus speeding up the development of the pipelines. Area experts and collaborators would switch between the streams to provide necessary technical expertise.

Communication during the hackathon was facilitated by Slack integration with Trello (for tasks management and progress tracking) and code developed was pushed to GitHub (for live code integration). Table 2 lists the various communication media used during the hackathon. Some groups also utilized Google docs for documenting their progress prior to migrating documentation into GitHub README files.

Remote participation in the hackathon was facilitated through the MConf conference system. One stream had a participant with very strong coding skills working remotely from the US; who managed to make progress on the corresponding workflow when the other group members were not working due to the big time difference between the USA and South Africa (SA). This ensured continuous development on the workflow when the team in SA would clock off and provide a to-do list which was accomplished by the participant from the US. Noticeable during the hackathon was the team spirit created and the increasingly later end time for the days (with most days ending at 8:30 pm as participants continued working after the different streams provided their daily reports). All participants wished for an extra day or two to complete their pipelines.

Post-hackathon activities

After the week-long hackathon at the University of Pretoria, members of each stream continued working on their respective pipelines communicating via Slack and Trello. Meetings were held over MConf every two weeks to report on the progress of each pipeline. Upon completion, each group handed their pipeline to other groups to test on different platforms, and thereby avoid bias in implementation and improve the documentation. Consequently, this facilitated the use of the four pipelines developed within H3Africa projects as highlighted in Table 1.

Discussion

The H3ABioNet Cloud Computing Hackathon was aimed at producing portable, cloud-deployable Docker containers for a variety of bioinformatics workflows including variant calling, 16S rDNA diversity analysis, quality control, genotype calling, and imputation and phasing for genome-wide association studies. The workflows developed in this hackathon benefited from workflow management systems, and further come with Docker recipe files that can be used to build container images when downloading images might be an issue. Thus, Dockerization provided a method to package and manage software, tools and workflows within a portable environment/container, similar to virtualization but with a smaller computing overhead compared to virtualization

The novelty of the H3ABioNet Cloud Computing Hackathon was that all the participants selected were involved in the latter stages of the planning and the setting of some of the outcomes for the hackathon. Critical recommendations during the hackathon planning meetings were that the resulting Docker containers and pipelines developed should be compatible with heterogeneous African research compute environments with portability and good documentation being key. This is especially important considering the fact that access to Cloud computing environments within Africa is still in its infancy. Hence, it was decided that development and testing of the pipelines should occur on a single machine, with the ability to be ported to a cluster or an HPC environment, and ultimately tested and deployed on cloud-based platforms (Amazon, Microsoft Azure, EGI FedCloud, IBM Bluemix, and the new African Research Cloud initiative).

Besides contributing solutions to African problems, three factors contributed to the success of this highly ICT-based activity in an African setting: 1) Almost all the communications tools used (Table 2) had equivalent apps that work right off a smartphone, a feature that many people within Africa (and less developed countries) tend to make use of 17. 2) The used tools were complementary to each other, and integration was sought whenever possible (like between Slack and Trello). 3) The hackathon was timed at the end of the 4th year of the initial H3ABi-oNet round of funding. At that point, the consortium (via its Infrastructure Working group) had already invested in improving the computational infrastructure within the network 11, including tools for regular communications and webinars 18. In a sense, Table 2 also represents our vetted list of collaborative tools in the light of 4 years of feedback from the consortium.

Lessons learnt and concluding remarks

The opportunity to link people physically and focus solely on one project has been highly effective in providing the main outline and proof of concept outputs. However, once people were back home, continuing the tasks has been a challenge. Clearly defining the roles and commitment of all the participants in the papers reporting the results should encourage them to complete the work, and increase their accountability.

The communication and management tools used for this hackathon (Table 2) were important as these tools facilitated interaction between and across team members and enabled the participants to continue to work in a structured manner once back at their respective institutions, despite time zones differences.

The H3ABioNet Cloud Computing Hackathon has been an important milestone for the Network as it brought together people with various skills to work on focused projects. It signalled the shift from capacity building to utilizing the capacity developed in order to tackle problems specific to the heterogeneous African computing environments, as defined and implemented by the mostly African participants. Equally important, this hackathon was not done in isolation from the rest of the scientific community nor could it have succeeded without local collaborations. This aspect, i.e. welcoming input and actively seeking it when needed from outside the consortium, is key to truly empowering the local community.

As software packages and computing environments evolve with varying build cycles and new bioinformatics tools become available, we envision that hackathons to keep these pipelines current, adopt new technology implementations such as Singularity, and develop new workflows such as for RNA-Seq analysis will occur. The pipelines developed during the H3ABioNet Cloud Computing hackathon will be used for training and data analyses for intermediate level bioinformatics workshops, and for scientific collaborations requiring bioinformatics expertise for data analysis such as with the H3Africa genotyping chip and GWAS analyses. Future H3ABioNet hackathons would also provide an opportunity to utilize the skills of trained bioinformaticians at intermediate and advanced levels, who would not otherwise attend bioinformatics training workshops, to come together to derive practical solutions that are of benefit to the African and wider scientific community.

Data and software availability

All data underlying the results are available as part of the article and no additional source data are required.

The four pipelines are available publicly via H3ABioNet's GitHub organization page https://github.com/h3abionet in the

following repositories: (h3agwas, chipimputation, h3agatk and h3abionet16S) as well as container images on quay.io at quay.io/organization/h3abionet_org and dockstore at: https://dockstore.org/workflows/h3abionet/h3agatk

All code is available under MIT license.

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Juan Ruiz-Alzola 🗓



Institute for Biomedical and Health Research, University of Las Palmas de Gran Canaria, Las Palmas de Gran Canaria, Spain

The authors have improved their original submission. In my view the article is now ready to be approved.

Competing Interests: No competing interests were disclosed.

Reviewer Expertise: Medical imaging, medical technology for sustainable development, cooperation for development

I confirm that I have read this submission and believe that I have an appropriate level of expertise to confirm that it is of an acceptable scientific standard.

Reviewer Report 04 September 2019

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Steffen Möller 🔟



Institute for Biostatistics and Informatics in Medicine and Ageing Research, Rostock University Medical Center, Rostock, Germany

Well done!

Competing Interests: No competing interests were disclosed.

I confirm that I have read this submission and believe that I have an appropriate level of expertise to confirm that it is of an acceptable scientific standard.

Version 1

Reviewer Report 22 June 2018

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Summary and impression

The article describes the organization and execution of a hackathon in Africa with the goal of producing cloud-deployable Docker containers for four bioinformatic workflows.

The article begins with a brief history of H3ABioNet and the need for bioinformatics infrastructure, training, and community in Africa.

Then the authors describe their pre-hackathon preparations, the weeks of activities during the hackathon, and their lessons learnt. No quantitative assessment of the hackathon was provided, but the authors do provide links to the bioinformatic workflows used.

Hackathons are an increasingly popular way of building tools and community for research and education. This article nicely describes why a hackathon was needed in Africa and how the authors went about organizing and executing their first hackathon. The article contains some useful suggestions for what bioinformatic or social tools can be used to facilitate research and communication. I think it is important to communicate this type of activity to the broader scientific community, however, I have a few concerns.

Major issue

My main concern is that this paper was submitted as a "method article". My understanding is that methods articles should describe new empirical or computational methods that are described in sufficient detail such that they can be reproduced. While this article does describe the authors' strategy for organizing and running a hackathon, this article is written more as a retrospective piece on the event rather than a recipe for running a hackathon. I am not convinced that it should be published as a method paper; unfortunately, there doesn't appear to be a more suitable platform within the AAS journal. https://aasopenresearch.org/for-authors/article-guidelines/method-articles

Minor issues

1. The figure legend does include a title, but it does not include a description of the key points nor does it explain the meaning of the arrows. According to the AAS guidelines, "the legend should be

sufficiently detailed so that it can stand alone from the main text". Additionally, it is not clear from first glance that "4 portable computational workflows" is the goal of the hackathon. This could be made more clear.

- 2. Table 1 provides a nice overview of communication channels. Can you elaborate and add what tools you used for sharing documents (e.g notes, slides, pdf)?
- 3. On page 4, the authors state: "Vital in setting up the teams...". Does this paragraph refer to the expertise of the learners/participants or to the people who are leading the stream or both?
- 4. On page 5, the first sentence of the discussion provides the first clear statement of the aim of the hackathon (in my opinion). The goal is mentioned in the abstract and intro, but it isn't as clear. It is in this paragraph that I realized that you had working pipelines, but they were not "dockerized". After reading this, the figure made a lot more sense. I recommend revising the abstract and intro to make it clear what the starting point (5 bioinformatic workflows not in the cloud) and the endpoints (4 bioinformatic workflows in the cloud).
- 5. The paragraph on "Post-hackathon feedback and actions" seems incomplete or perhaps is mislabelled. This paragraph describes communication and work that extended past the hackathon, but it does not describe any assessment or feedback mechanisms that were used. Also, what happened after groups traded platforms? Was the documentation improved or was this simply the goal?
- 6. The article jumps from "Introduction" to "Discussion" without providing a clear a description of some of the results or outcomes. Is there a reason why some statistics regarding participation or progress toward the goal are not reported? If there is a reason why demographic information cannot be provided, that is fine, but a report about how much progress was made toward the goal of dockerization would be useful.
- 7. While I appreciate that you provided a link to the GitHub repo and Docker container, I highly recommend getting a DOI for these repositories. The AAS data guidelines page provides a list of providers. Some of your GitHub repositories already have versions, so it should be fairly easy to import these into Zenodo for a DOI. https://aasopenresearch.org/for-authors/data-guidelines.

Is the rationale for developing the new method (or application) clearly explained? Partly

Is the description of the method technically sound?
Partly

Are sufficient details provided to allow replication of the method development and its use by others?

Partly

If any results are presented, are all the source data underlying the results available to ensure full reproducibility?

No source data required

Are the conclusions about the method and its performance adequately supported by the findings presented in the article?

Partly

Competing Interests: No competing interests were disclosed.

We confirm that we have read this submission and believe that we have an appropriate level of expertise to confirm that it is of an acceptable scientific standard, however we have significant reservations, as outlined above.

Reviewer Report 02 May 2018

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Institute for Biomedical and Health Research, University of Las Palmas de Gran Canaria, Las Palmas de Gran Canaria, Spain

I have overall enjoyed reading this paper. The implementation of the hackathon is well explained, and provides good hints for others to replicate both in Africa and in the rest of the World. In particular, a large array of ICT-based collaborative tools has been applied whose use is very interesting everywhere. The motivation is also very relevant, as hands-on knowledge sharing is important to build distributed strong scientific communities that can develop new solutions and keep collaboration active within wide multi-site networks. This is particularly so in the case of bioinformatics, a profoundly interdisciplinary and complex field, key to modern knowledge-based economy, health and welfare, where Africa could leverage important opportunities as far as some hurdles are properly managed. It is in this framework where I see the main contribution of this paper, and I must point out the relevance of the collaboration among African and rest of the World institutions and researchers, in addition to the internal African collaboration. In my view all of this is essential to unlock all the potential, and achieve not only scientific success but a wider interaction of great value for economic development. I'm not getting into any technical detail of the pipelines since, as reported in the paper, they will be presented elsewhere.

Nevertheless I have some constructive criticisms that I'd like to share:

- 1. I've pointed out that the rationale is only partly explained essentially because some boundary conditions should be explained: how many nodes and researchers are participating, what their access to Internet is like in their sites, what difficulties they're experiencing in their home institutions to access to knowledge and to interact with other international colleagues, etc. Africa is large with very different situations across countries and within each country. It'd be good that the reader could understand better the everyday situation that motivates an action like this, or a wider program such as H3ABioNet. It'd also be good to explain a bit longer what H3A and H3ABioNet are, and how they become important programs for the scientific development of Africa.
- 2. The description of the method is technically sound, but I have some concerns about how realistic it is to expect such sophisticated ICT-based collaboration, difficult anywhere, considering the

existing difficulties for high bandwidth connections in many areas of Africa, including many Universities in main cities. Our group is currently involved in a cooperation with research groups in four African countries. Internet access, good enough for sharing medical datasets, for example, or just to hold videoconferences, is something that we cannot take for granted. Again, this is also related to (1) and boundary conditions, but it is my experience that dealing with this sort of issues is key for success and, at least, to replicate the experience elsewhere.

3. The conclusions are somehow conditioned by what I've already pointed out.

In summary, I think this is a very interesting and motivating paper that could benefit of a better explanation of the boundary conditions, and of how the specific difficulties have been overcome, so that others can replicate the experience in the same or in different fields across the continent.

Is the rationale for developing the new method (or application) clearly explained? Partly

Is the description of the method technically sound?

Partly

Are sufficient details provided to allow replication of the method development and its use by others?

Yes

If any results are presented, are all the source data underlying the results available to ensure full reproducibility?

Yes

Are the conclusions about the method and its performance adequately supported by the findings presented in the article?

Partly

Competing Interests: No competing interests were disclosed.

Reviewer Expertise: Medical imaging, medical technology for sustainable development, cooperation for development

I confirm that I have read this submission and believe that I have an appropriate level of expertise to confirm that it is of an acceptable scientific standard, however I have significant reservations, as outlined above.

Reviewer Report 18 April 2018

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Steffen Möller (1)



Institute for Biostatistics and Informatics in Medicine and Ageing Research, Rostock University Medical Center, Rostock, Germany

The paper describes an in-person meeting in Africa to further develop workflows for genomic analyses and distribute that skill throughout the continent. As a Northern European I can only remotely assess the difficulties of computational biology and bioinformatics services in Africa in local areas. I can see how important such Hackathons are to edutRain research groups with difficulties to attend international conferences. And, with some experience in attending and organizing such events, I was very curious about what may be different in Africa. After all, in Europe there are communities having difficulties to commute, too. The European Union has extra funds for graduate students from Eastern European countries for instance. But there are also highly talented high school students who are under age or do not have the funds or scientific contacts to travel/be trained. So, if you have developed principles for African talents to overcome such obstacles to learn about such Hackathons and prepare for them, then I would be very much interested to see what we can adopt over here.

From what I then read in the article, there was not so much that the authors did differently, except that they seem to have done it particularly well. Table 1 describes a whole bunch of communication channels when typically it is a mere Wiki or co-editing site orchestrating the participants. Still, every attendee had to physically attend, except that the event was in Africa describing an African set of resources and there was external expertise flown in. I tend to think that here the event fell short of what could have been (and is often) done to invite remote participation. Also, one could videograph training material to support the further distribution of these Africa-specific analysis skills.

I am also a bit critical about the dominance of online resources that demand good internet connections as in the download of gigabytes of data: Docker. Here I wish more would be done to promote offline services. But I am biased, the authors NM and MRC know me as a contributor to Debian Med, MRC being a Debian contributor himself, which basically means that participants could take a DVD home and perform analyses of their samples without Internet access involving as many local-to-them machines as they like. For the workflows one doesn't need most of the complicated bits for which one needs computational expertise the article is describing. And that may have helped the post-Hackathon drop in participation, while, of course every event sees that drop and that is a main reason to have such a dedicated time to jointly develop our research environments in the first place.

Concerning the scientific results, I understand that there is a separate paper prepared. Still, can you say as much as if there are scientific papers out there that already employed the pipelines for their analysis? I mean, from the time before the Hackathon? This would emphasize that you are indeed redistributing a very current set of skills with practical acceptance in the community.

There is something else to it all. Hackathons form a social network of trust. And you need trust in locally well-described samples. The genetic diversity of Africa is a gem, but one needs to be aware of the demands for population stratification. And because of the harsh environment conditions, one can expect considerable batch effects on samples taken, for which the emphasis on equally collected control samples is important. You can certainly read about it all, but it will help to hear in person about that study that was ruined because cases and controls were kept separate and one box had the dry ice evaporated early – factor variation and confounding technical parameters cannot be communicated enough. Well-established workflows allow participants to analyse their local data independently, have extra parameters matching local concerns, with matching local controls, which will all improve the quality of the pan-African study at large.

So, I do not think that anything performed for the organisation of this Hackathon was specific to Africa. In the contrary, there should have been some teleconferencing to it. The "initial day to get everyone up to speed" reminds me a bit more of a Summer School than a Hackathon, for which it is not uncommon that most participants already know each other for some time and would not need that day. Could you possible elude a bit more on how the work performed was structured? And how was the external expertise intermingled with your local needs for Africa?

I would want to retitle the paper towards something like "Hackathon on workflows for genomics held in Africa". The H3ABioNet workflows are nicely accepted everywhere, I tend to think, so, why not have an H3ABioNet workflow Hackathon in Europe with some Africans flying in? Clarifying that a bit more may help the paper.

Is the rationale for developing the new method (or application) clearly explained? Partly

Is the description of the method technically sound? Partly

Are sufficient details provided to allow replication of the method development and its use by others?

Yes

If any results are presented, are all the source data underlying the results available to ensure full reproducibility?

Yes

Are the conclusions about the method and its performance adequately supported by the findings presented in the article?

Partly

Competing Interests: No competing interests were disclosed.

I confirm that I have read this submission and believe that I have an appropriate level of expertise to confirm that it is of an acceptable scientific standard, however I have significant reservations, as outlined above.

Author Response 21 Apr 2018

Victor Jongeneel, University of Illinois at Urbana-Champaign, Champaign, USA

Thanks very much to Steffen Möller for his pointed comments. It is certainly true that this hackathon did not differ in any major way from similar events held in high-income countries, and did not incorporate any features specific to the African context. Its primary value was to expose African scientists to the practical aspects of community development of computer code, and to try to create a community around the maintenance of a set of workflows that implement methods that are useful to the H3Africa research community and beyond.

There are aspects of the work that may not have come across in the paper. For one thing, the

workflows implementing haplotyping, imputation, and GWAS analysis were based on work done in the framework of the H3Africa AWIGen project, and are in production for the analysis of data generated by this project. Similarly, the workflow for variant calling from WGS data was used in the analysis of 350 African genomes that has led to the design of a novel genotyping chip optimized for African populations, and the 16S rDNA sequence analysis was derived from work done to analyze bacterial populations present in leg ulcers of sickle cell patients in Nigeria. Therefore, all of the code developed during the hackathon was solidly anchored in existing genomic analysis projects in Africa.

Secondly, the workflows developed in the hackathon serve as practical implementations of Standard Operating Procedures for the H3Africa Accreditation Exercises, which are used to evaluate the capacity of African research groups to analyze complex genomic datasets being produced by its research projects (see Jongeneel et al, PLoS Comput Biol, PMC5453403). Success in taking one of the exercises is considered a landmark for African groups who are preparing to step into the existing gap between data production and data analysis, where the analysis is typically undertaken by First World groups.

It is true that the authors, including myself, could have done a better job at explaining how the hackathon and its products are anchored in the H3Africa research ecosystem. I hope that the above clarifies this.

As a final remark, while highly Internet-dependent tools were used extensively during the hackathon, to my knowledge none required a very high bandwidth. At least two of the participants attended remotely from North America, and were able to contribute substantially in part because of their time zone differences and asynchronous contributions.

Competing Interests: No competing interests were disclosed.

Reader Comment 04 May 2018

Steffen Möller,

Thank you for your constructive reply to my initial comments. This addresses all my reservations and I am looking forward for a revised upload. SM

Competing Interests: No competing interests were disclosed.