National ethics guidance in Sub-Saharan Africa on the collection and use of human biological specimens: a systematic review

Francis Barchi 1* and Madison T. Little 2

Abstract

**Background:** Ethical and regulatory guidance on the collection and use of human biospecimens (HBS) for research forms an essential component of national health systems in Sub-Saharan Africa (SSA), where rapid advances in genetic- and genomic-based technologies are fueling clinical trials involving HBS and the establishment of large-scale biobanks.

**Methods:** An extensive multi-level search for publicly available ethics regulatory guidance was conducted for each SSA country. A second review documented active trials listed in the WHO International Clinical Trials Registry Platform as of January 2015 in which HBS collection was specified in the protocol. Findings were combined to determine the extent to which countries that are study sites for HBS-related research are supported by regulatory guidance language on the collection, use, ownership and storage of biospecimens.

**Results:** Of the 49 SSA countries, 29 had some form of national ethics guidance, yet only 17 provided language relating to HBS-related research, with specific guidance on consent (14), ownership (6), reuse (10), storage (9), and export/import/transfer (13). Ten countries accounted for 84% of the active clinical trials involving the collection of HBS in SSA. All except one of these countries were found to have some national guidance in the form of regulations, codes of ethics, and/or standard operating procedures; however, only seven of the ten offered any language specific to HBS.

**Conclusions:** Despite the fact that the bulk of registered clinical trials in SSA involving HBS, as well as existing and proposed sites for biorepositories under the H3Africa Initiative, are currently situated in countries with the most complete ethics and regulatory guidance, variability in the regulations themselves may create challenges for planned and future pan-African collaborations and may require legislative action at the national level to revise. Countries in SSA that still lack regulatory guidance on HBS will require extensive health system strengthening in ethics governance before they can be full participants in the modern research enterprise.

**Keywords:** Sub-Saharan Africa, Biospecimens, Biobanks, Ethics guidance, Regulatory systems, Ethics committees, Health systems, Materials transfer agreements

Background

Exponential growth in the fields of pharmacogenetics and genomics research over the past decade is fueling worldwide interest in the establishment of biorepositories of human biological specimens. Biorepositories already exist in three SSA countries—The Gambia, South Africa, Zimbabwe [1–3]. Major expansion in this domain is occurring in Sub-Saharan Africa, largely driven by the Human Heredity and Health in Africa Initiative (H3Africa), a joint effort of the National Institutes of Health (NIH), the Wellcome Trust, and the African Society for Human Genetics [4]. The ‘hub and spoke’ model adopted by this initiative calls for the development of four pilot regional biobanks – two in South Africa and one each in Nigeria and Uganda – which will serve as multi-country repositories of specimens collected under sponsored research activities taking place throughout the region [5]. The H3Africa Initiative reflects significant efforts by the international community of researchers,
funding agencies, and African academic institutions to build a foundation for sophisticated, state-of-the-art genomic research on burdensome diseases in Africa; the ultimate goal of the initiative is to improve the health of the African peoples.

From its inception, the initiative has sought broad consensus on topics relating to ethics and governance. A Joint NIH and Wellcome Trust Policy document sets forth high-level principles on ethics, governance, and resource sharing with which research networks and programs funded by the initiative are expected to comply [6]. A standing working group on ethics and regulatory issues comprising ethicists and other members of the scientific community meets periodically to discuss and develop recommendations on a wide range of topics, including community engagement strategies, cultural issues, informed consent, participant safety, return of results, intellectual property, material transfer agreements, and other regulatory issues. While such guidance has the potential to serve as a model framework for all countries in Africa to consider as they develop their own national guidelines, the current focus of the initiative is on those entities that it supports. Applied more broadly, the pan-continental approach espoused by H3Africa and similar initiatives may require a level of ethical guidance at the country level explicitly addressing the collection and use of HBS which is currently not to be found in most national ethics regulatory guidance. In many African countries, such systems are in their infancy; policies that address emerging technologies such as whole genome sequencing and association studies are absent [7, 8].

The International Declaration on Human Genetic Data, promulgated by UNESCO in 2003, recognized the importance of national policies to guide the collection, processing, use, and storage of human genetic data, as well as the review by local and institutional ethics committees of protocols involving such data [9]. In Africa specifically, a growing body of literature has focused on the need to have ethical and legal governance structures in place to oversee research involving HBS and the operation of biobanks [10, 11].

If the H3Africa vision of a pan-continental genomic enterprise is to be fully realized, ethics guidelines at the national level must align in ways that support interoperability among African countries. In cases where no guidelines presently exist, investment in institutional capacity-building by such entities as H3Africa and others may be needed to support their development. In cases where national guidelines do exist, some may require the addition of new language specific to research involving HBS; such revisions may require legislative approval. There may be instances where national values simply will not support the transnational transfer of genetic information, broad data sharing within research communities (however relevant to their populations), or the delegation of decision-making on future use to an outside authority. The lack of interoperability among African nations in their ethics guidance as well as differences in interpretation of meaning in instances where international recommendations have been adopted has the potential to slow, even derail, the realization of benefit from new advances and approaches in biomedical research [10, 12].

Although a number of scholars have examined the ethical, legal, and social issues associated with the collection and use of biospecimens in research, there has been far less focus on the existence and nature of regulatory guidance on HBS at the national level in Sub-Saharan Africa [8, 11–17]. Of particular importance is the identification of national ‘gaps’ in ethics guidance, as well as inconsistencies and national differences among those countries where guidance does exist.

**Methods**

This study involved two systematic reviews designed to determine the availability of publicly accessible regulatory HBS guidance at the national level in SSA countries and to assess the extent to which it addressed ethical and regulatory issues relating to consent, ownership, reuse, storage, and export/import/transfer. Two reviews were conducted as part of this project. The first involved a multi-level search to identify publicly available national research ethics guidance on the collection and use of HBS in countries in Sub-Saharan Africa. This process, detailed in Table 1, involved keyword searches, on-line reviews of international ethics databases and regional and national government websites, and, in those instances where on-line efforts yielded no results, direct contact via email with national ethics committees and scans of ethics review statements included in previously published journal. National ethics guidance was operationalized to include enabling legislation for the creation and constitutions of national research institutes and national research ethics committees, codes of ethics, national ethics guidelines, and standard operating procedures. We excluded those documents that only provided instructions to researchers on how to submit their protocols, instruction on writing consent documents, and organizational information from research ethics committees that were not identified as being national in jurisdiction. Materials Transfer Agreements (MTA) and Specimen Transfer Agreements (STA) for specific countries were included in our search but were only used as a source of guidance if we could find a specific reference to “human” biological specimens. We limited our results to those national documents that we could access publicly on-line or secure through contact with the national regulatory body, reasoning that guidelines that were inaccessible via such mechanisms would be of limited utility to the research community.
A second review was conducted using the WHO International Clinical Trials Registry Platform (ICTRP) to identify registered clinical trials taking place in Sub-Saharan Africa as of January 1, 2015, and, of these, the number of clinical trials in each country that called for the collection of HBS as part of the study protocol [18]. The ICTRP is a web-based, publicly accessible compilation of studies that have been registered in any of the following databases: Australian New Zealand Clinical Trials Registry (ANZCTR); Chinese Clinical Trial Registry (ChiCTR); ClinicalTrials.gov; EU Clinical Trials Register (EU-CTR); ISRCTN; The Netherlands National Trial Register (NTR); Brazilian Clinical Trials Registry (ReBec); Clinical Trials Registry – India (CTRI); Clinical Research Information Service – Republic of Korea (CRiS); Cuban Public Registry of Clinical Trials (RPCEC); German Clinical Trials Register (DRKS); Iranian Registry of Clinical Trials (IRCT); Japan Primary Registries Network (JPRN); Pan African Clinical Trial Registry (PACTR); Sri Lanka Clinical Trials Registry (SLCTR); and Thai Clinical Trials Register (TCTR). Included in the count of active clinical trials were those trials listed in the database as active/not recruiting; recruiting; not yet recruiting; authorized-recruitment/may be ongoing or finished; and pending. Studies were excluded if identified as not recruiting; complete; terminated; closed/follow-up continuing; closed to recruitment/ follow-up complete; temporary halt or suspension; other; not applicable; or withdrawn. The process used to identify active trials calling for HBS collection is illustrated in Fig. 1.

Results
Table 2 summarizes the availability of national ethics guidance irrespective of content as well as guidance language specific to HBS in each of the 49 countries...
| Country                      | Ethics system guidance (Genl) | Specific HBS language | Ethics system guidance specific to HBS | HBS informed consent | HBS ownership | HBS reuse | HBS storage | HBS export import transfer | HBS materials or specimens transfer agreement |
|-----------------------------|--------------------------------|-----------------------|---------------------------------------|----------------------|---------------|-----------|-------------|---------------------------|---------------------------------------------|
| Angola                      | ●                              | ●                     |                                       |                      |               |           |             |                           |                                             |
| Benin                       | ●                              | ●                     |                                       |                      |               |           |             |                           |                                             |
| Botswana                    | ●                              | ●                     | ●                                    | ●                    | ●             | ●         | ●           | ●                         | ●                                           |
| Burkina Faso                | ●                              | ●                     | ●                                    | ●                    |               |           |             |                           |                                             |
| Burundi                     | ●                              |                       |                                       |                      |               |           |             |                           |                                             |
| Cameroon                    | ●                              |                       |                                       |                      |               |           |             |                           |                                             |
| Cape Verde                  | ●                              |                       |                                       |                      |               |           |             |                           |                                             |
| Central African Rep.        | ●                              |                       |                                       |                      |               |           |             |                           |                                             |
| Chad                        | ●                              |                       |                                       |                      |               |           |             |                           |                                             |
| Comoros                     | ●                              |                       |                                       |                      |               |           |             |                           |                                             |
| Rep. of Congo               | ●                              |                       |                                       |                      |               |           |             |                           |                                             |
| Congo (DRC)                 | ●                              |                       |                                       |                      |               |           |             |                           |                                             |
| Cote d'Ivoire               | ●                              |                       |                                       |                      |               |           |             |                           |                                             |
| Djibouti                    | ●                              |                       |                                       |                      |               |           |             |                           |                                             |
| Equatorial Guinea           | ●                              |                       |                                       |                      |               |           |             |                           |                                             |
| Eritrea                     | ●                              |                       |                                       |                      |               |           |             |                           |                                             |
| Ethiopia                    | ●                              | ●                     |                                       |                      |               |           |             |                           |                                             |
| Gabon                       | ●                              |                       |                                       |                      |               |           |             |                           |                                             |
| The Gambia                  | ●                              | ●                     | ●                                    | ●                    | ●             | ●         | ●           | ●                         | ●                                           |
| Ghana                       | ●                              |                       |                                       |                      |               |           |             |                           |                                             |
| Rep. of Guinea              | ●                              |                       |                                       |                      |               |           |             |                           |                                             |
| Guinea-Bissau               | ●                              |                       |                                       |                      |               |           |             |                           |                                             |
| Kenya                       | ●                              | ●                     | ●                                    | ●                    |               |           |             |                           |                                             |
| Lesotho                     | ●                              |                       |                                       |                      |               |           |             |                           |                                             |
| Liberia                     | ●                              | ●                     | ●                                    | ●                    |               |           |             |                           |                                             |
| Madagascar                  | ●                              |                       |                                       |                      |               |           |             |                           |                                             |
| Malawi                      | ●                              | ●                     | ●                                    | ●                    |               |           |             |                           |                                             |
| Mali                        | ●                              |                       |                                       |                      |               |           |             |                           |                                             |
| Mauritania                  | ●                              |                       |                                       |                      |               |           |             |                           |                                             |
| Mauritius                   | ●                              |                       |                                       |                      |               |           |             |                           |                                             |
| Mozambique                  | ●                              |                       |                                       |                      |               |           |             |                           |                                             |
| Namibia                     | ●                              |                       |                                       |                      |               |           |             |                           |                                             |
| Niger                       | ●                              | ●                     | ●                                    | ●                    |               |           |             |                           |                                             |
| Nigeria                     | ●                              | ●                     | ●                                    | ●                    |               |           |             |                           |                                             |
| Rwanda                      | ●                              | ●                     | ●                                    | ●                    |               |           |             |                           |                                             |
| Sao Tome & Principe         | ●                              | ●                     | ●                                    | ●                    |               |           |             |                           |                                             |
| Senegal                     | ●                              |                       |                                       |                      |               |           |             |                           |                                             |
| Seychelles                  | ●                              |                       |                                       |                      |               |           |             |                           |                                             |
| Sierra Leone                | ●                              | ●                     | ●                                    | ●                    |               |           |             |                           |                                             |
| Somalia                     | ●                              | ●                     | ●                                    | ●                    |               |           |             |                           |                                             |
| South Africa                | ●                              | ●                     | ●                                    | ●                    |               |           |             |                           |                                             |
included in the WHO region constituting Sub-Saharan Africa. Detailed information on the specific content of any HBS-related language is provided in Table 3. Twenty-nine (60 %) of the countries in the region had some national ethics guidance, either in the form of laws, regulations, codes, guidelines, or standard operating procedures. Of these 29 countries, however, twelve did not have regulations that addressed specific guiding principles or rules for ethics research or review. Six (Congo DRC, Equatorial Guinea, Republic of Guinea, Lesotho, Madagascar, and Mali) had regulatory language that was limited to establishing or defining the functions and constitution of research ethics committees. One (Mauritius) had drafted a code of research ethics that was distributed for public comment in 2003, but no documentation could be found that it had been operationalized. Six (Congo DRC, Equatorial Guinea, Republic of Guinea, Lesotho, Madagascar, and Mali) had regulatory language that was limited to establishing or defining the functions and constitution of research ethics committees. One (Mauritius) had drafted a code of research ethics that was distributed for public comment in 2003, but no documentation could be found that it had been operationalized. Three countries (Cameroon, Mozambique, Rwanda) offered limited language specific to guiding principles or ethical conduct of research, but link their review processes to the ethical guidance contained in the Declaration of Helsinki, CIOMS, or the ICH-GCP [19–21]. No national guidelines for general ethical conduct of research could be found for The Gambia, but the Medical Research Council – The Gambia had published very detailed guidelines for the collection and use of biospecimens stored in the National DNA Bank of The Gambia which we have included in our analysis [22]. In addition, two countries (Chadand Gabon) had been working with UNESCO on the establishment of national ethics committees but no regulatory documentation could be found [23, 24].

More than one half (59 %) of the countries whose guidelines articulated ethical principles or guidelines for ethical review include some language to guide the collection and use of HBS for biomedical research. Fourteen of these offered some language with respect to informed consent; 10 addressed the subject of reuse; and 12 contained specific language relating to export/import/transfer. Only six countries offered language with respect to HBS ownership and nine contained language regarding storage. Thirteen required that a Materials or Specimens Transfer Agreement be executed. Table 4 reports the findings with respect to registered, active clinical trials in Sub-Saharan Africa that called for the collection of HBS. Of the 5319 studies identified in the WHO International Clinical Trials Registry Platform, 1802 of were currently active as of January 1, 2015. Three hundred sixty-six (366) of these involved the collection of a biosample from research subjects. The distribution of these trials across the region are illustrated in Fig. 2.

Table 5 reports on the ethics system guidance on HBS found in the countries where the largest number of studies involving HBS are currently taking place in Sub-Saharan Africa. These 10 countries account for nearly 82 % of the active clinical trials involving the collection of HBS in the region that were registered in the WHO International Clinical Trials Registry Platform as of January 1, 2015. They included the five countries that either have existing biorepositories (The Gambia, South Africa and Zimbabwe) or planned to establish regional biorepositories in the near future as part of the H3Africa Initiative (Nigeria, South Africa and Uganda). All ten of these countries had some form of ethics system guidance in the form of regulations, codes of ethics, standard operating procedures, or guidelines, and, with the exception of Mali, provided some regulatory language at the national level as to the collection and use of HBS. Notably only four of these countries (The Gambia, Kenya, Uganda, and Zambia) included language in their national ethics documents on the ownership of HBS. Mali, where at the time of our study there were currently 10 registered active trials requiring a biosample, had no guidance language on HBS in its national ethics documents. Although most of the countries in which large biorepositories existed or are planned as part of H3-Africa contain fairly detailed HBS language in their guidelines, only two (The

**Table 2** SSA countries with HBS-related information in legislative/regulatory/guidance ethics documents (Continued)

| South Sudan | Sudan | Swaziland | Tanzania | Togo | Uganda | Zambia | Zimbabwe |
|-------------|-------|-----------|----------|------|--------|--------|----------|
| ●           |       | ●●       | ●        |      | ●●●●   | ●●●●   | ●●●●●●   |
| TOTAL       | 29    | 17        | 14       | 6    | 10     | 9      | 12       | 13  |
| Country       | Legislation, regulations, and/or guidance on collection and use of HBS | Specific HBS language | HBS consent | HBS ownership | HBS secondary use | HBS storage | HBS Import/export |
|--------------|------------------------------------------------------------------------|-----------------------|-------------|---------------|------------------|-------------|------------------|
| Angola       | None found                                                            | None found            | None found  | None found    | None found       | None found  | None found       |
| Benin        | Loi 2005-31 Portant prévention, prise en charge et contrôle du VIH/SIDA (2006) [28] | Loi 2005-31 portant prévention, prise en charge et contrôle du VIH/SIDA en République du Benin (2005) | No language | No language   | No language      | No language | No language      |
| Botswana     | Ministry of Health Standard Operating Procedures, Article 7.2 Procedures on Human Genetic Research, pages 103-108. Articles 8.1-8.5, Human Biological Materials, pages 111-114 | Consent required unless waived by an ethics committee.* (MOH SOPs, 2011, 7.2.iv) An ethics committee may sometimes waive, with or without conditions, the requirement for consent. (MOH SOPs, 2011, 7.2.v) Acquisition, storage and future use of biological materials: Informed consent process and forms separate from and in addition to informed consent for research participation. (MOH SOPs, 2011, 8.2) | Government of Botswana is normally the owner of research data. The creators of the data, the Principal Investigators and local institutions in Botswana should hold the data in trust on behalf of the Botswana government and the research participant. The host institutions or researcher has to apply to the HRDC regarding the decisions about use and sharing of their data with other researchers and institutions according to Botswana law (MOH SOPs, 2011, 8.5, Appendix 11).* Research protocols should detail the purpose and use of research specimens Where archived specimens are required in another experiment, researchers must provide a new protocol for review and approval by HRDC. (MOH SOPs, 2011, 8.2.iii) Host institution in Botswana is entrusted custodianship of samples and should hold samples in trust on behalf of Botswana Government and the research participant. Research subjects have right to withdraw their samples if traceable. Health Research and Development Committee approval required for use, transfer, storage & future use according to Botswana laws. Where samples have not been obtained as part of research, the institution that collected them takes custodianship of them. Any future study on such samples is subject to review by HRDC. (MOH SOPs, 2011, 8.2) | No transfer unless: a) the researcher and the other research group are collaborating on research that has been approved by an ethics committee; and b) the genetic material cannot be identified. Transfer of identifiable or potentially identifiable genetic material may be approved by HRDC in certain circumstances. (MOH SOPs, 2011, 7.2.vi) All exchanges and transfers (including importation) of HBS for research purposes shall require clearance from the HRDC. (MOH SOPs, 2011, 8.3) Procedures for Exchange/Transfer of HB Materials, pg. 112 (MOH SOPs, 2011, 8.3) |
**Table 3** National regulatory guidance language on collection and use of HBS by country* (Continued)

| Legislation, regulations, and/or guidance re. research and/or HBS | Specific HBS language | HBS consent | HBS ownership | HBS secondary use | HBS storage | HBS Import/export |
|---|---|---|---|---|---|---|
| Burkina Faso | Law No. 23/94/ADP of 19 May 1994 on the Code of Public Health [34] | Human biological materials means the organs, tissues and products that can be used for therapeutic or research purposes on human beings. They can be derived from living or cadaveric donors. (Law No. 23/94) | Consent required for human genetic research unless waived by an ethics committee. Institutions or organizations wishing to conduct research on genetic material and on information collected for non-research purposes should develop and disseminate a general policy that informs subjects that such material and information may be used for future research, following research ethics committee approval, participants in such institutions or organizations should be informed that this policy exists and that their privacy and confidentiality will be protected. (p. 105-106). | No language | No language | No language | No language |

| Burundi | None found | None found | None found | None found | None found | None found | None found |
| Cameroon | Arrêté No. 079/A/MSP/DS-Order of the Minister of Public Health, October 22, 1987: Creation and organization of an ethics committee for research involving human subjects [37] | No language | No language | No language | No language | No language | No language |
| Legislation, regulations, and/or guidance re. research and/or HBS | Specific HBS language | HBS consent | HBS ownership | HBS secondary use | HBS storage | HBS Import/export |
|---|---|---|---|---|---|---|

(Article 9 refers to an evaluation process relying on Declaration of Helsinki)
Decision 0674/D/MSP/CIRCMB, October 13, 2006:
Creation and functions of an ethics committee in the Chantal International Reference Center for research on the prevention and treatment of HIV/AIDS [38]

Cape Verde
None found

Central African Republic
None found

Chad
Part of UNESCO Bioethics project. Efforts to establish National Ethics Commission reported in meeting report, UNESCO, 2008) [23]

Comoros
None found

Republic of Congo
None found

Congo (DRC)
Ministry of Health, Politique nationale de recherché sur les systems de santé en RDC (June 2004) [39]

Côte D’Ivoire
None found

Djibouti
None found

Equatorial Guinea
None found
Table 3 National regulatory guidance language on collection and use of HBS by country* (Continued)

| Legislation, regulations, and/or guidance re. research and/or HBS | Specific HBS language | HBS consent | HBS ownership | HBS secondary use | HBS storage | HBS Import/export |
|---------------------------------------------------------------|-----------------------|-------------|---------------|------------------|-------------|------------------|
| Presidential Decree No. 124/2014: Establishes the National Ethics Committee of Equatorial Guinea (CENGIE), 2014. [40] Presidential Decree 125/2014: Constitute CENGIE and appoints inaugural members, 2014 [41] | No language | No language | No language | No language | No language | No language |
| Eritrea | None found | None found | None found | None found | None found | None found |
| Ethiopia | None found | None found | None found | None found | None found | None found |
| National Health Research Ethics Review Guideline [5th Ed.] Sept. 2014 [42] | National Health Research Ethics Review Guideline, Guideline 8.3.2, p. 51-52 | IC process should include info on HBS and data to be collected, data anticipated to be derived from sample as well as health and other records to be accessed, their intended uses, storage and duration of storage. Participants must be informed that specific information will not be shared with family members, and that, if shared with third parties, will be anonymized. Participant should be given option of allowing sharing of samples or not. (Guideline, 8.3.2, p. 52) Separate consent form for HBS collected for storage and/or future use. Participant may choose whether to have HBS stored for future studies. (Guideline, 9.2, p. 57) | Host institution in Ethiopia holds HBS in trust on behalf of research participant. (Guideline, 9-2, p. 58) Where HBS have not been obtained as part of research, the institution that collected the samples takes custodianship. (Guideline 9.2, p. 58) | Any shared samples must be anonymized to the recipient, Use of HBS beyond what is stated in original protocol must have consent of research participants or their representatives. Secondary use may only be done on anonymized samples and after getting approval by the IRB. When subsequent use of HBS or data is proposed that is not consistent with original I/C, a new consent should be obtained from participant/guardian or LAR or a waiver of consent should be requested from IRB. (Guideline, 8.3.2, p. 52) | Intent to store HBS and place and duration of storage must be specified in initial consent document (Guideline, 8.3.2) Research participants reserve right to withdraw their HBS from storage if samples are linked. (Guideline, 9.2) | MTA required. It must contain details regarding purpose of transfer/export, arrangements for collaboration and benefit-sharing, a framework for accessing and sharing data, restrictions to third party transfers, and annual reports to host institution. An Ethiopian scientist must be included as a co-investigator in all future studies using the HBS. The IRB in Ethiopia must review all research studies on stored HBS. (Guideline 9.2, p. 58) |
| Gabon | No language | No language | No language | No language | No language | No language |
| Legislation, regulations, and/or guidance re. research and/or HBS | Specific HBS language | HBS consent | HBS ownership | HBS secondary use | HBS storage | HBS Import/export |
|---|---|---|---|---|---|---|
| The Gambia (Republic of) | Guidelines for The Gambia National DNA Bank, 2001 [22] | Samples from children only after permission from parents or guardians. No consent for reuse, but permission of Gambian government/MRC Laboratories Ethics Committee is required. Permission from donor for reuse of samples that have not been anonymized is required in principle, but committee can waive this requirement. (DNA Bank Guidelines, 2001) | DNA Collections are jointly owned by the Gambian people through their government representatives and the Medical Research Council (DNA Bank Guidelines, 2001) | Reuse permissible with approval from Joint Committee. Different levels of permissions/consents required depending on whether or not requested specimens have been anonymized, collected as part of a specific protocol or available to researchers with approval from Committee only. [DNA Bank Guidelines, 2001] | The Gambia National DNA Bank was established in 2001, the first biobank on the African continent (DNA Bank Guidelines, 2001). | Permission for export based on signed agreement between researchers and the DNA Bank, with approval of Committee. Agreement must describe analyses to be undertaken with specimens; feedback must be given to MRC Labs, results of analyses must eventually be placed in the public domain, and any manuscripts resulting from research must be submitted to Joint Committee prior to submission. (DNA Bank Guidelines, 2001) |
| Ghana | No research ethics regulatory guidance Public Health Act 2012 [Act 851] Good Clinical Practice Doc No FDA/SMC/CTD/GL- GCP/2013/02 Ver. 2, 18 Dec 2015 (Adopted 27 Jan 2016) [45] | No language | No language | No language | No language | No language |
| Guinea (Republic of) | Decree No. D/218/PRG/SGG: On the Establishment, Functions and Organization of the National Ethics Committee for Research in Health (CNERCS) (Decree No D/218). 1998. [46] Decree contains no ethics regulatory guidance language. | No language | No language | No language | No language | No language |
| Guinea-Bissau | None found | None found | None found | None found | None found | None found |
| Country     | Legislation, regulations re. specific HBS language | Consent | Ownership | Secondary use | Storage | Import/export |
|------------|-----------------------------------------------|--------|----------|--------------|--------|---------------|
| Kenya      | No language                                   | No language | No language | No language | No language | No language |
| Lesotho    | No language                                   | No language | No language | No language | No language | No language |
| Legislation, regulations, and/or guidance re. research and/or HBS | Specific HBS language | HBS consent | HBS ownership | HBS secondary use | HBS storage | HBS Import/export |
|---------------------------------------------------------------|----------------------|------------|---------------|------------------|------------|-----------------|
| **Liberia**<br>Ethics approval required from one of three REC bodies: Liberia Institute of Biomedical Research/National Health Science Research EC (LIBR/NHSREC), the University of Liberia Institutional Review Board (UL-IRB), or the newly established National Research Ethics Board (NREB). [52]<br>University of Liberia-Pacific Institute for Research and Evaluation: Institutional Review Board (IRB) Policies and Procedures Handbook 2008 [53] Liberia Medicines and Health Products Regulatory Authority, Guideline for Application to Conduct Clinical Trials in Liberia 2014. [54] | Research involving the collection or study of pathological specimens, or diagnostic specimens may be exempt, if those sources are publicly available or if the information is recorded by the investigator in such a manner that the subjects cannot be identified, directly or indirectly, through identifiers linked to the subjects. Archival research in which individual subjects could potentially be identified is not exempt. (University of Liberia, Policies & Procedures Handbook, 2008, Art. 5 pg. 23) | No language | No language | No language | No language | Materials Transfer Agreement must be provided to the Liberia Medicines and Health Products Regulatory Authority’s (LMHRA) (LMHRA Guidelines, 2014.) |
| **Madagascar**<br>Decree No. 5855/99-SAN (June 17, 1999) modified by Decree No. 4583/2000-SAN (May 8, 2000) (Created a National Ethical Committee for Biomedical Research.) Documents contain no research ethics regulatory guidance. [55] | No language | No language | No language | No language | No language | No language |
| Country   | Specific HBS Language                                                                 | HBS Consent | HBS Ownership | HBS Secondary Use | HBS Storage | HBS Import/Export |
|-----------|---------------------------------------------------------------------------------------|-------------|---------------|------------------|-------------|------------------|
| Malawi    | Clear explanation and justification for the collection and exporting of biological samples will have to be made. (General Guidelines, 2007, Section 6.1) | Informed consent for HBS required. Persons may only be consented for HBS collection for purposes of answering the study objectives of a presently intended study that has been clearly defined. (Policy Requirements, 2007, 10.0) | No language   | All forms of studies & testing aimed at collecting and storing HBS for future unspecified genetic research/analysis, including scientific retrospective analysis is non-permissible (HBS Guidelines, 2012, 3.4.7) | Procedures for specimen storage will have to be clearly defined. (General Guidelines, 2007, Section 6.1) | Analysis of specimens should be done within Malawi by local technicians/professionals. Export permitted only in exceptional circumstances when the needed technology does not exist in Malawi nor can it be imported or when tests are needed to confirm results and/or quality control and validation are required. (Policy Requirements, 2007, 10.0) | MTAs are required. Agreement must clearly specify why export is necessary, its intended use, length of time HBS will be kept, and name of local technician/ professional who will be responsible for HBS testing. (Policy Requirements, 2007, 10.0) |
| Mali      | No language                                                                           | No language | No language   | No language      | No language | No language      |
| Organization and functions of the National Institute of Public Health Research, including establishment of an ethics committee within Institute | Mauritania |
|---|---|
| Legislation, regulations, and guidance re. research and/or HBS | None found |
| Specific HBS language | None found |
| HBS consent | None found |
| HBS ownership | None found |
| HBS secondary use | None found |
| HBS storage | None found |
| HBS import/export | None found |
| Legislation, regulations, and/or guidance re. research and/or HBS | Specific HBS language | HBS consent | HBS ownership | HBS secondary use | HBS storage | HBS Import/export |
|-------------------------------------------------------------|----------------------|-------------|---------------|------------------|-------------|------------------|
| Namibia                                                      | None found           | None found  | None found    | None found       | None found  | None found       |
| Ministry of Health and Social Sciences has constituted a Biomedical Research Ethical Committee but it is not operational. [64] |                      |             |               |                  |             |                  |
| Niger                                                        | None found           | None found  | None found    | None found       | None found  | None found       |
| Nigeria                                                     | None found           | None found  | None found    | None found       | None found  | None found       |
| National Health Research Ethics Committee of Nigeria (NHREC) National Code of Health Research Ethics 2007. [65] Policy Statement on Storage of Human Samples in Biobanks and Biorepositories in Nigeria (PS1.02013) (November 1, 2013) [66] | Reference to HBS in connection with MTAs (Ethics Code, 2007) See Biobanks (Biobank Policy, 2013) NHREC to provide sole oversight of ethical aspects of biobanking (Biobank Policy, 2013) | Biobanks in Nigeria must retain copies of all consent forms and these must be matched to all samples in biobanks (Section D.3.i.a., Biobanks, 2013) NHREC supports Broad Consent- consent in which the type or purpose of research is defined in broad terms and for a work that is not specified by time (Section E.1, Biobanks, 2013) | No language | No language | Any samples kept more than 2 months post-analysis are considered 'banked' and covered by Biobank Policy. Copies of all participant I/C forms must be available and can be matched to samples stored in biobanks (Biobank Policy 2013) | Export permitted, subject to MTA approved by HREC. Institutional HRECs shall grant final approval for research involving international transfer of HBS. (HREC Code, 2007) |
| Rwanda                                                       | Rwanda Ministry of Health, National Research Ethics Committee: Standard Operating Procedures, 2009 [67] Rwanda Ministry of Health, National Health Research Committee: Operational Guidelines, 2012. [68] | Regulations with respect to HBS (Sections 35 & 36 of Rwanda Ministry of Health Standard Operating Procedures, 2009. | HBS must be collected with free and informed consent even if tissue is obtained as part of patient care. Consent forms must indicate if HBS is being collected for current research only, how long the specimens will be kept, and when they will be destroyed. If HBS is stored longer than the current research, then a separate permission for storage must be obtained. | Under MTAs: Ownership is the "Provider of the samples." Ownership of any income resulting from commercialization must be negotiated in good faith (Rwanda SOPs, 2009, Appendix 7). | No language | No language | MTAs are required. Entities to whom HBS is transferred must provide any resultant publications, and provider must be part of the publication team. (Rwanda SOPs, 2009, Appendix 7) |
| Legislation, regulations, and/or guidance re. research and/or HBS | Specific HBS language | HBS consent | HBS ownership | HBS secondary use | HBS storage | HBS Import/export |
|---------------------------------------------------------------|------------------------|-------------|---------------|------------------|------------|-----------------|
| Sao Tome & Principe                                           | None found             | None found  | None found    | None found       | None found | None found       |
| Senegal                                                       |                        | No language | No language   | No language      | No language | No language      |
| Arrêté ministerial No. 3224 MSP: DERF-DER, March 17, 2004: Creation and organization of National Council for Health Research (CNRS) [69] Loi No. 2009-17, March 9, 2009: Code of Ethics for Health Research. [70] Règlement Intérieur du Conseil National de la Recherche en Santé [Standard Operating Procedures], March 7, 2006. [71] | Section 3.9 Biological specimens/samples (Guidelines for conducting clinical trials, 2014, p. 20-21) Definition of HBS: a biological specimen or a biological sample is defined as material derived from various animal and human sources (e.g., blood, tissues, and cells) used to treat and prevent diseases. (Guidelines for GCP, 2014) | Separate consent for use of HBS (Guidelines, 2014, Sec. 3.9.1) HBS taken in course of clinical care may be used for research without consent subject to ethics committee review. Patients have right to know their HBS is being used for research. Patient refusal to such use does not warrant waiver of consent. Refusals to be honored except in case of public health emergency. (Guidelines, 2014, Sec. 3.9.2) | No language | No language | No language |
| Seychelles                                                    | None found             | None found  | None found    | None found       | None found | None found       |
| Sierra Leone                                                  | Pharmacy Board of Sierra Leone: Guidelines for Conducting Clinical Trials of Medicines, Food Supplements, Vaccines, and Medical Devices in Sierra Leone (Version 01) (G-SLClinTrial), 2014. [72] Pharmacy Board of Sierra Leone: Guideline for Good Clinical Practice (GCP) in Sierra Leone (Version 01) (SL-GCPs). 2014 [7373] Ministry of Health & Sanitation, Office of the Sierra Leone Ethics and Scientific Review Committee: Sierra Leone Ethics and Scientific Review Committee – Guidelines. n.d.[74] | Secondary use of HBS is constrained by conditions specified in initial consent. Therefore, initial consent should specify whether or not there will be secondary use; conditions under which subjects must be re-contacted; plans, if any, for de-identification; and subject rights. (Guidelines, 2014, Sec. 3.9.3) | No language | No language | Materials Transfer Agreement is required. (NIAID Communication with the Pharmacy Board of Sierra Leone (PBSL) (October 2014–January 2015) |
| Legislation, regulations, and/or guidance re. research and/or HBS | Specific HBS language | HBS consent | HBS ownership | HBS secondary use | HBS storage | HBS Import/export |
|---------------------------------------------------------------|-----------------------|-------------|---------------|------------------|-------------|------------------|
| Somalia                                                       | None found            | None found  | None found    | None found       | None found  | None found       |
| Act 61 of 2003 National Health Act [75]                       | Chapter 8, Sections 53-68. (National Health Act, 2003) Regulations Relating to the Use of HBS (2012) | Written informed consent of individual with provisions for consent from subjects who are minors or mentally ill (Regulations, 2012, 3.1) Written informed consent by users/donors for release of stored information and for long-term storage of genetic material, stem cells, or research findings (Regulations, 2012, 13.d & f). | No language | HBS information used for purposes for which it was originally intended. (Regulations, 2012, 13.e). | Written informed consent of the user or donor for long-term storage of genetic material, stem cells, or research findings. HBS information treated confidentially. Users’ written informed consent prior to release of stored information. (Regulations 2012, 13e & f). | Export permit is required. No export unless it is established that sample was donated under terms of Act and will be used in accordance with terms of Act. (Act, 2003, 8.68.1(g)) |
| South Sudan                                                  | None found            | None found  | None found    | None found       | None found  | None found       |
| SUDAN                                                        | Guidelines, 2008, Secs. 5.2, 5.10, & 5.11. | Investigator must provide subjects: policy to use results of genetic tests and familial genetic information and precautions to prevent disclosure to others (Guidelines, 2008, 5.2.16); possible research sites, direct or secondary use of HBS taken in course of clinical care (5.2.18); disposal, storage, future use of HBS (5.2.19); any commercial products from HBS and distribution of any revenues (Guidelines, 2008, 5.2.20); Consent forms must have separate sections requesting use of HBS for research purposes (Guidelines, 2008, 5.10); HBS taken in course of clinical care may be used w/o consent subject to approval of ethics committee (Guidelines, 2008, 5.11). | No language | No language | No language |
| Country     | Legislation, regulations, and/or guidance re. research and/or HBS | Specific HBS language | HBS consent | HBS ownership | HBS secondary use | HBS storage | HBS Import/export |
|-------------|------------------------------------------------------------------|-----------------------|-------------|---------------|-------------------|-------------|------------------|
| Swaziland   | None found                                                       | None found            | None found  | None found    | None found        | None found  | None found       |
| Tanzania    | National Institute for Medical Research Act, 1979 [78]           | Part V Human DNA      | Consent for intended study only. New protocol required for reuse (Guidelines, 2009, 8.8) No collection of a sample without the consent of the sample source (DNA Act, 2009, 38-41). | Samples of Human DNA are the property of the sample source. (DNA Act, 2009, 28.2.b) Where the research results from an individual or an institute are new or unique, the researcher or the institute shall have the intellectual property rights (DNA Act, 2009, 5.3.7). | Consent for intended study only. New protocol required for reuse (8.8, Guidelines, 2009) Sample destroyed on completion of analysis unless sample source’s representative has previously directed otherwise in writing (DNA Act, 2009, 28.2.e) After completion of original research, investigators must (a) within one month notify of their intent to reuse and secure the permission of sample source or (b) destroy samples within three months (38.44, DNA Act, 2009, 38.44) | No language | Materials transfer agreement is required. |
| Togo        | None found                                                       | None found            | None found  | None found    | None found        | None found  | None found       |
| Uganda      | Uganda Ministry of Health, National Health Policy, 1999 [81]     | See Guidelines 2014, Sec. 10, pp. 28-31. | Consent must include explanation of how HBS will be managed at end of study. If stored, separate consent must be obtained, with purpose of study, quantities stored, location, measures to ensure confidentiality, future governance, risks and potential benefits. (Guideline, 2014, 5.3.h) | Sample sources own the samples if samples are identified, source may withdraw them at any time. Samples to be held in trust by duly authorized Uganda organization. Trustee organization has authority to decide use, transfer, storage and future use of HBS in its care, taking into account rights and welfare of research participants (Guidelines, 2014, 10.3) | None found | None found | None found |
|             | Uganda National Council for Science and Technology (NCST), National Guidelines for Research Involving Humans as Research Participants, July 2014, [82] | See HBS I/C           |             |               |                  |             |                  |

*HBS: Human Biological Samples*
Table 3 National regulatory guidance language on collection and use of HBS by country* (Continued)

| Legislation, regulations, and/or guidance re. research and/or HBS | Specific HBS language | HBS consent | HBS ownership | HBS secondary use | HBS storage | HBS Import/export |
|---------------------------------------------------------------|-----------------------|-------------|---------------|-----------------|------------|------------------|
| Zambia                                                        |                       |             |               |                 |            |                  |
| The National Health Research Act 2013 [83]                    | May only be collected for purposes stated in research protocol (Act, Part VI) | No HBS removed from living person for health research purposes without written consent of donor in accordance with provisions of the Act, Act, Art. 47 (1) | Ownership specified in MTA as determined by Minister of health in consultation with Health Authority. Act, Art. 51 (1)(a) | A person shall not withdraw HBS from a living person for any unspecified future health research activity of unspecified storage. Act, Art. 47 (2) | A person shall not withdraw HBS from a living person for any unspecified future health research activity or unspecified storage. Act, Art. 47 (2) | Only with written approval of the National Health Research Authority Act, Art. 50 (1) and if terms of MTA are met Art. 50 (2) MTA is required. Minister, in consultation with Authority, decides on requirements. |
| Scientific Technological Research Act (Research Act) (Ch. 10:22), 1986 (Replaced Research Act 1959) [84] | Discussion of HBS not included in MRCZ guidelines. | No language | No language | No language | Extraterritorial storage of biospecimens beyond Research Council of Zimbabwe approval period is illegal. RCZ STA, 2015) Note: All international studies must be registered with RCZ. | Export for specified research purposes only. No third party transfers. Approval of REC required. Specimens Transfer Agreement (STA) is required. |

* Guidance language cited verbatim where possible.
### Table 4: Clinical trials in SSA involving HBS collection (Source: WHO-ICTRP as of January 1, 2015)

| Country               | Total # studies | Total active recruiting/Not yet recruiting | Total active not recruiting | Total active studies | HBS collection active recruiting/Not yet recruiting | HBS collection active not recruiting | Total trials collecting HBS |
|-----------------------|-----------------|------------------------------------------|----------------------------|----------------------|---------------------------------|------------------------------------|-------------------------------|
| Angola                | 3               | 2                                        | 0                          | 2                    | 2                               | 0                                  | 2                             |
| Benin                 | 33              | 10                                       | 1                          | 11                   | 2                               | 1                                  | 3                             |
| Botswana              | 50              | 10                                       | 5                          | 15                   | 4                               | 1                                  | 5                             |
| Burkina Faso          | 117             | 20                                       | 11                         | 31                   | 3                               | 1                                  | 4                             |
| Burundi               | 0               | 0                                        | 0                          | 0                    | 0                               | 0                                  | 0                             |
| Cameroon              | 57              | 12                                       | 6                          | 18                   | 1                               | 1                                  | 2                             |
| Cape Verde            | 0               | 0                                        | 0                          | 0                    | 0                               | 0                                  | 0                             |
| Central African Rep.  | 6               | 5                                        | 3                          | 8                    | 0                               | 0                                  | 0                             |
| Chad                  | 0               | 0                                        | 0                          | 0                    | 0                               | 0                                  | 0                             |
| Comoros               | 0               | 0                                        | 0                          | 0                    | 0                               | 0                                  | 0                             |
| Republic of Congo     | 59              | 10                                       | 3                          | 13                   | 3                               | 0                                  | 3                             |
| Congo (DRC)           | 17              | 10                                       | 2                          | 12                   | 1                               | 0                                  | 1                             |
| Cote d'Ivoire         | 37              | 10                                       | 4                          | 14                   | 3                               | 0                                  | 3                             |
| Djibouti              | 0               | 0                                        | 0                          | 0                    | 0                               | 0                                  | 0                             |
| Equatorial Guinea     | 0               | 0                                        | 0                          | 0                    | 0                               | 0                                  | 0                             |
| Eritrea               | 1               | 0                                        | 0                          | 0                    | 0                               | 0                                  | 0                             |
| Ethiopia              | 74              | 22                                       | 7                          | 29                   | 2                               | 3                                  | 5                             |
| Gabon                 | 34              | 5                                        | 0                          | 5                    | 1                               | 0                                  | 1                             |
| The Gambia            | 79              | 11                                       | 4                          | 15                   | 5                               | 1                                  | 6                             |
| Ghana                 | 138             | 33                                       | 7                          | 40                   | 10                              | 1                                  | 11                            |
| Guinea                | 74              | 1                                        | 2                          | 3                    | 0                               | 0                                  | 0                             |
| Guinea-Bissau         | 43              | 5                                        | 9                          | 14                   | 2                               | 5                                  | 7                             |
| Kenya                 | 320             | 65                                       | 27                         | 92                   | 17                              | 4                                  | 21                            |
| Lesotho               | 0               | 0                                        | 0                          | 0                    | 0                               | 0                                  | 0                             |
| Liberia               | 7               | 1                                        | 1                          | 2                    | 0                               | 0                                  | 0                             |
| Madagascar            | 12              | 1                                        | 0                          | 1                    | 1                               | 0                                  | 1                             |
| Malawi                | 171             | 33                                       | 15                         | 48                   | 7                               | 4                                  | 11                            |
| Mali                  | 101             | 20                                       | 7                          | 27                   | 7                               | 3                                  | 10                            |
| Mauritania            | 5               | 3                                        | 0                          | 3                    | 0                               | 0                                  | 0                             |
| Mauritius             | 9               | 2                                        | 1                          | 3                    | 0                               | 0                                  | 0                             |
| Mozambique            | 52              | 11                                       | 1                          | 12                   | 2                               | 1                                  | 3                             |
| Namibia               | 0               | 0                                        | 0                          | 0                    | 0                               | 0                                  | 0                             |
| Niger                 | 19              | 7                                        | 1                          | 8                    | 2                               | 0                                  | 2                             |
| Nigeria               | 103             | 37                                       | 7                          | 44                   | 10                              | 2                                  | 12                            |
| Rwanda                | 57              | 8                                        | 1                          | 9                    | 2                               | 0                                  | 2                             |
| Sao Tome & Principe   | 0               | 0                                        | 0                          | 0                    | 0                               | 0                                  | 0                             |
| Senegal               | 58              | 10                                       | 3                          | 13                   | 2                               | 0                                  | 2                             |
| Seychelles            | 2               | 0                                        | 1                          | 1                    | 0                               | 1                                  | 1                             |
| Sierra Leone          | 11              | 3                                        | 3                          | 6                    | 1                               | 2                                  | 3                             |
| Somalia               | n.d.            | n.d.                                      | n.d.                       | n.d.                 | n.d.                            | n.d.                               | n.d.                          |
| South Africa          | 2712            | 832                                      | 234                        | 1066                 | 150                             | 39                                 | 189                           |
Discussion

This study examined the availability of national ethics and regulatory guidance on biomedical research in Sub-Saharan African countries and identified the extent to which national regulatory systems provided ethics guidance on specific aspects of HBS collection and use to inform research design and ethics review. Ethical principles and regulatory guidance regarding HBS consent, ownership, reuse, storage, and export/import/transfer are foundational elements in national health systems as research activities calling for the collection and use of biospecimens in Africa expand and ethics committees are increasingly called on to review and respond to rapidly advancing innovation in research and research technologies.

Our research found that despite efforts at the international, regional, and national levels, 20 of the countries in Sub-Saharan Africa (41%) at the time of our study still lacked articulated national ethical principles and regulatory guidance for policy-making, review, and monitoring of research. Among those countries where ethical and regulatory guidance existed, specific language with respect to HBS collection and use was often lacking or incomplete. While in 17 of the countries in Sub-Saharan Africa

Table 4 Clinical trials in SSA involving HBS collection (Source: WHO-ICTRP as of January 1, 2015) (Continued)

| Country      | No data | 0   | 1   | 2   | 3   | 4   | 5   |
|--------------|---------|-----|-----|-----|-----|-----|-----|
| South Sudan  | 0       | 0   | 0   | 0   | 0   | 0   | 0   |
| Sudan        | 0       | 0   | 0   | 0   | 0   | 0   | 0   |
| Swaziland    | 7       | 4   | 0   | 4   | 1   | 0   | 1   |
| Tanzania     | 263     | 45  | 14  | 59  | 10  | 3   | 13  |
| Togo         | 8       | 5   | 0   | 5   | 0   | 0   | 0   |
| Uganda       | 351     | 74  | 25  | 99  | 20  | 4   | 24  |
| Zambia       | 147     | 31  | 11  | 42  | 5   | 4   | 9   |
| Zimbabwe     | 82      | 23  | 5   | 28  | 7   | 2   | 9   |
| Total        | 5319    | 1381| 421 | 1802| 283 | 83  | 366 |

No data (n.d.)

Gambia and Uganda) spoke to issues of ownership. After Mali, Zimbabwe offered the least guidance within this group of countries, with no language on HBS-specific consent, ownership, or reuse.

Fig. 2 Distribution of active clinical trials in SSA involving HBS collection (Trial information Source: WHO-ICTRP as of January 1, 2015; Map source: Wikimedia Commons)
(59%), some form of national regulatory guidance relating to HBS did exist, a number of these lacked explicit guidance on HBS-related issues such as consent (3), ownership (11), reuse (7), storage (8), and export/import/transfer (5).

Gaps in national ethical and regulatory guidance in the region on the collection and use of HBS for research may result in inconsistent reviews within ethics committees and pose challenges for full participation by African countries in genomic research. In the absence of guiding principles and regulatory direction, ethics committees may delay or deny HBS-related research of potential benefit to their communities, or they may approve such research without due exploration of its implications for its citizenry or its cultural values.

Differences in ethical and regulatory requirements among African countries where guidance does exist also appear to raise impediments to the interoperable pan-African approach to genomic research espoused by H3Africa. Despite the fact that the bulk of registered clinical trials involving HBS in Sub-Saharan Africa, as well as existing sites and those sites proposed for future biorepositories, are currently situated in countries with the most complete ethics and regulatory guidance on HBS, variability in the regulations themselves may create challenges for pan-African collaborations that rely on transnational sharing of specimens and data or that delegate decision-making responsibility with respect to future use to repository-centric ethics or scientific review committees. Malawi, for example, contained language in its guidelines that all forms of studies and testing aimed at collecting and storing HBS for future unspecified genetic research/analysis is ‘non-permissible’ [25]. Tanzania required that a new consent be secured for each new research protocol in which a sample will be reused [26]. Uganda, where one of the H3Africa repositories is intended to be located, required that all requests for transfers of HBS be made by a legal resident of Uganda who was affiliated with a locally registered and recognized Ugandan organization [27].

While these and other potential sources of regulatory tension may be resolved in the future, for now they represent potential constraints on research that calls for a more ‘regional’ perspective of specimen and data sharing, and are likely to require legislative action at the national level to revise. At the same time, gaps in HBS-related guidance within countries that are already research-intensive sites may give rise to lengthy or inconsistent review decisions by local ethics committees left to make their own policy decisions in the absence of clearly stated regulations or principles at the national level.

Future growth in the fields of pharmacogenetics and genomic research is likely to generate demand for HBS from a more representative pool of the African peoples and to prompt researchers to look for new communities in countries where research is currently not active. The transnational movements of diseases themselves, as seen in the 2014 Ebola outbreak in Western Africa, may identify new target populations for study beyond those of interest at present. Such expansion will require system-wide capacity-building in these settings and will need to be guided by clearly articulated national regulatory guidance to ensure ethical and thoughtful research governance.

Resources to assist national governments in developing ethics capacity may be available through such programs as UNESCO’s Assisting Bioethics Committees (ABC) program, which aims to build and strengthen bioethics infrastructure within its member states and which is already working to build capacity in Chad, Gabon, and Madagascar [23, 24, 28]. H3Africa’s working group on ethics and regulatory issues and its work to date in articulating broad principles and regulatory guidance regarding HBS collection and use [6] would, were its focus to extend beyond its own funded

Table 5 National ethics guidance on 10 countries host to the most registered clinical trials

| Country      | # Active studies w/HBS | Existing biobanks | H3Africa regional biobanks | Ethics system guidance (Gen'l) | Ethics system HBS language | Ethics system guidance specific to HBS |
|--------------|------------------------|-------------------|-----------------------------|-------------------------------|---------------------------|---------------------------------------|
| South Africa | 189                    | ●●●●●●●●●●        | ●●●●●●●●●●                  | ●                            | ●●●●●●●●●●                  | ●●●●●●●●●●                                  |
| Uganda       | 24                     | ●●●●●●●●●●        | ●●●●●●●●●●                  | ●                            | ●●●●●●●●●●                  | ●●●●●●●●●●                                  |
| Kenya        | 21                     | ●●●●●●●●●●        | ●●●●●●●●●●                  | ●                            | ●●●●●●●●●●                  | ●●●●●●●●●●                                  |
| Tanzania     | 13                     | ●●●●●●●●●●        | ●●●●●●●●●●                  | ●                            | ●●●●●●●●●●                  | ●●●●●●●●●●                                  |
| Nigeria      | 12                     | ●●●●●●●●●●        | ●●●●●●●●●●                  | ●                            | ●●●●●●●●●●                  | ●●●●●●●●●●                                  |
| The Gambia   | 11                     | ●●●●●●●●●●        | ●●●●●●●●●●                  | ●                            | ●●●●●●●●●●                  | ●●●●●●●●●●                                  |
| Malawi       | 11                     | ●●●●●●●●●●        | ●●●●●●●●●●                  | ●                            | ●●●●●●●●●●                  | ●●●●●●●●●●                                  |
| Mali         | 10                     | ●●●●●●●●●●        | ●●●●●●●●●●                  | ●                            | ●●●●●●●●●●                  | ●●●●●●●●●●                                  |
| Zambia       | 9                      | ●●●●●●●●●●        | ●●●●●●●●●●                  | ●                            | ●●●●●●●●●●                  | ●●●●●●●●●●                                  |
| Zimbabwe     | 9                      | ●●●●●●●●●●        | ●●●●●●●●●●                  | ●                            | ●●●●●●●●●●                  | ●●●●●●●●●●                                  |
| Total        | 309                    | 3                 | 4                           | 10                           | 9                         | 8 4 7 8 8 8                                  |
research networks and programs, be a valuable starting point for countries as they begin to build national consensus on the value of genomics research for their communities and the region as a whole.

**Limitations of this study**
Data used in this study were largely derived from publicly accessible sources. Despite an extensive search, some of the countries in Sub-Saharan Africa that we report as lacking national ethics regulatory guidelines may, in fact, have them. Given the central role that national policies play in shaping the research landscape in the region, it is important that researchers and ethics committees have ready access to country-level guidance on the collection and use of HBS as the scope of genomics research in Sub-Saharan Africa expands. In addition, the mapping of active clinical trials that involve HBS in Sub-Saharan Africa relied on publicly accessible international registries. It therefore did not take into account those studies which are not required to register, and serves more as an illustration of a potential regulatory ‘gap’ between research and national governance than as a comprehensive source of data on HBS-related research in the region.

**Conclusion**
Despite substantial capacity-building efforts, many countries still lack regulatory guidance on the collection and use of human biological specimens in research. Although the African countries currently most in demand as HBS-related clinical research sites number among those offering the most guidance, extensive health system strengthening will be needed to ensure that regulatory language is available to guide the ethical extension of HBS-related research into other countries in the region. Efforts to create interoperability across national policies to meet international and regional research goals of Pan-African research collaboration must be on-going, recognizing that some differences may be value-based choices by representative governments while others, although malleable to change, will require legislative action to amend.

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**Availability of data and materials**
All data relevant to the analysis of regulatory information is accessible using the links provided in the references for this manuscript. All data used in the analysis of current clinical trials in Africa registered on clinicaltrials.gov are available through that web site.

**Authors’ contributions**
FB conceptualized the project on which this manuscript is based, designed the methodology, participated in data collection, analyzed the findings, and drafted the original manuscript. MTL had the lead in data collection, assisted in the organization of findings, and read and commented on draft versions of the manuscript. Both authors read and approved the final version of the manuscript.

**Competing interests**
The authors declare that they have no competing interests.

**Consent for publication**
All authors consent to the publication of this manuscript.

**Ethics approval and consent to participate**
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**Author details**
1Edward J. Bloustein School of Planning and Public Policy, Rutgers University-New Brunswick, 33 Livingston Street, New Brunswick, NJ 08901, USA.
2Department of Social Policy & Intervention, University of Oxford, Oxford, UK.

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