## NOTE #8

ZIKA in Brazil Real Time Analysis (ZiBRA–2): an RRI experience

By Marta Giovanetti, Fernanda Khouri, Luiz Alcantara

# **RRI IMPLEMENTATION** IN BIOSCIENCE ORGANISATIONS

GUIDELINES FROM THE STARBIOS2 PROJECT

Andrea Declich with the STARBIOS2 partners



STARBIOS2 project has received funding from the European Union's Horizon 2020 research and innovation programme under grant agreement No 709517.

### **NOTE #8**

ZIKA in Brazil Real Time Analysis (ZiBRA-2): an RRI experience

#### By Marta Giovanetti, Fernanda Khouri, Luiz Alcantara

Advances in DNA sequencing technology have ushered in a new era of pan-genomics and genomic surveillance, in which traditional molecular diagnostics and genotyping methods are being enhanced and even replaced by genomics-based methods to aid epidemiologic investigations of communicable diseases (Gardy et al., 2018). The ability to compare and analyse entire pathogen's genomes has allowed unprecedented resolution into how and why infectious diseases spread. The rapid development of these technologies has made sequencing of viral genomes possible and even routine (Shendure et al., 2008).

There are currently two major ways in which high-throughput sequencing technologies are used in public health and diagnostic applications, (i) to track outbreaks and epidemics in order to call for public health responses and (ii) to characterize individual infections to tailor treatment decisions (Theze et al., 2018; Faria et al., 2017). Focusing on these aims, genome sequencing has been successfully used to describe unique and detailed insights into the transmission, biology, and epidemiology of many health care-associated viral pathogens.

Considering the improvements on portability and quality of sequencing, and the acceleration and standardization of analytical pipelines, the applicable routine of genome sequencing may soon become the common *de facto* method for infectious disease control. In the context of virus investigations, pan-genomics and bioinformatics in general face great challenges. Rapid extraction of genomic features with an evolutionary signal facilitates

evolutionary analyses ranging from the reconstruction of species phylogenies to tracing epidemic outbreaks.

In February 2016, the World Health Organisation declared a Public Health Emergency of International Concern in response to the transmission of ZIKV in the Americas. In that context, the ZiBRA-2 project was launched as a multicentre collaboration between the University of Oxford, University of Birmingham, Evandro Chagas Institute, University of São Paulo and Oswaldo Cruz Foundation employing a promising approach to generating a substantial number of complete genome sequences for Zika virus (ZIKV) through MinION in a mobile laboratory trip.

The ZiBRA-2 project is based on principles of ethics, social engagement and open access to the information obtained. We consider that it is necessary to present the ZIKV results to other scientific communities and try to increase the participation of the public and civil society in bioscience research. Thus, during the project all the sequences and information generated are published in real-time on the ZiBRA-2 websites (http://www.zibraproject.org; https://www.zibra2project.org), and the final results are made available to society through scientific publications in open access journals.

Based on a previous genomic surveillance trip during the Ebola outbreak in Guinea in 2014-2015, the ZiBRA-2 project aimed to generate a large number of ZIKV complete genome sequences from the Northeast of Brazil covering a broad geographical region including historical samples, and from patients with a range of clinical presentations. The method consisted of genome-tiling PCR to enrich ZIKV material in clinical samples followed by library preparation prior to MinION loading (Faria et al., 2016; Quick et al., 2017).

The ZiBRA-2 team working together with the Central Laboratory of Public Health (LACEN) personnel, tested 1349 clinical samples for ZIKV RNA across Rio Grande do Norte, Paraíba, Recife, Maceió, and Bahia states and captured 850 mosquitoes from urban

and peri-urban fields in each place along the trip. The project also involved capacity building as each local team was trained to perform the whole protocol on subsequent trips. It is important to note that the team is composed of men and women who participate from the design of the study until the final publication and are trained at all stages, to reduce the gender discrimination. (Faria et al., 2016).

After the original trip that took place in June 2016, the ZiBRA-2 project has been extended and up to now trained teams to track not only ZIKV, but also other arboviruses circulating in Brazil including emerging and re-emerging strains. The team was also employed to investigate the dispersion of the CHIKV - East-Central South African genotype spreading in North Brazil (Naveca et al., 2019) as well as to characterize the largest Yellow Fever outbreak registered in Southeast Brazil in December 2016. By analysing 64 new yellow fever virus genomes the virus transmission pattern was revealed to originate in non-human primates, rejecting the hypothesis of urban transmissions.

As the mobile trips occur, more people are being trained to continue performing genomic surveillance throughout the country and also in some places in Africa like Angola and Cabo Verde (Hill et al., 2019). Also, the productivity of these trips is increasing each time, with generation of around 60 complete genome sequences in five days. Besides that, the development of faster protocols and more than 12 barcodes per run suggests this number will increase soon. A single flow cell used in MinION can run up to 96 genomes and produces reads up to 200 Kb in length, with a throughput of 1.5 Gb, and more than 100,000 reads at a single run. Ongoing improvements to the launched barcoding kits in the nanopore sequencing technology had the potential to increase the number of generated genomes per sequencing run from 12 to 96, which could also increase the number of genome sequences derived from affected regions and allow more detailed investigations of the

association between pathogen mutations and environmental context with less costs.

The participation of ZiBRA-2 in STARBIOS2 provides an ideal environment to showcase these research projects, and highlights the practice of Responsible Research and Innovation (RRI) in the context of this unique bioscience endeavour.

# REFERENCES

- Ahuja, S.K., Aiuti, F., Berkhout, B., Biberfeld, P., Burton, D.R., Colizzi, V., Deeks, S.G., Desrosiers, R.C., Dierich, M.P., Doms, R.W., Emerman, M., Gallo, R.C., Girard, M., Greene, W.C., Hoxie, J.A., Hunter, E., Klein, G., Korber, B., Kuritzkes, D.R., Lederman, M.M., Malim, M.H., Marx, P.A., McCune, J.M., McMichael, A., Miller, C., Miller, V., Montagnier, L., Montefiori, D.C., Moore, J.P., Nixon, D.F., Overbaugh, J., Pauza, C.D., Richman, D.D., Saag, M.S., Sattentau, Q., Schooley, R.T., Shattock, R., Shaw, G.M., Stevenson, M., Trkola, A., Wainberg, M.A., Weiss, R.A., Wolinsky, S., Zack, J.A. (2006). A plea for justice for jailed medical workers. Science, 314(5801).
- Alberts, B., Kirschner, M.W., Tilghman, S., &Varmus, H. (2014). Rescuing US biomedical research from its sistemi flaws. *Proceedings of the National Academy of Sciences*, 111(16).
- Alsop, R., Bertelsen, M., Holland, J. (2006). Empowerment in practice: from analysis to implementation, The International Bank for Reconstruction and Development/The World Bank.
- Ambrosio, A.M., Mariani, M.A., Maiza, A.S., Gamboa, G.S., Fossa, S.E., Bottale, A.J. (2018). Protocol for the production of a vaccine against Argentine Hemorrhagic Fever in Maria S. Salvato (ed.) Hemorrhagic Fever Viruses: Methods and Protocols. Methods in Molecular Biology, vol. 1604. Springer. Doi 10.1007/978-1-4939-6981-4\_24.
- Andoh, C.T. (2011). Bioethics and the challenges to its growth in Africa. *Open journal of philosophy.* **1**(02), 67-75. 10.4236/ojpp.2011.12012.
- Bamgbose, A. (2011). African languages today: The challenge of and prospects for empowerment under globalization. *In Selected proceedings of the 40th Annual Conference on African Linguistics*. ed. Eyamba G. Bokamba, et al., 1-14. Cascadilla Proceedings Project Somerville. www.lingref.com, document #2561.
- Barugahare, J. (2018). African bioethics: methodological doubts and insights. **BMC** *medical ethics*. 19(1), 98. 10.1186/s12910-018-0338-6.
- Battilana, J., Leca B., and Boxenbaum E. (2009). "How actors change institutions: towards a theory of institutional entrepreneurship." Academy of Management annals 3.1 (2009).
- BBSRC Biotechnology and Biological Sciences Research Council (2018). "Forward look for UK Bioscience" released on 29 September 2018. https://bbsrc.ukri.org/documents/forward-look-for-uk-bioscience-pdf/ (accessed on: 05/09/2019).
- Beckert, J. (1999). Agency, entrepreneurs, and institutional change. The role of strategic choice and institutionalized practices in organisations. Organisation studies, 20(5).
- Bendels, M.H., Dietz, M.C., Brüggmann, D., Oremek, G.M., Schöffel, N., Groneberg, D.A. (2018). Gender disparities in high-quality dermatology research : a descriptive bibliometric study on scientific authorships. *BMJ Open* 2018;8:1–11. doi:10.1136/bmjopen-2017-020089

- Berger, P., & Luckmann, T. (1969). La costruzione sociale della realtà. Il Mulino, Bologna.
- Besley, J.C., & Nisbet, M.C. (2013). How scientists view the public, the media and the political process. Public Understanding of Science, 22(6), 644–659. https://doi.org/10.1177/0963662511418743
- Bijker, W.E. & d'Andrea, L. (eds.) (2009). Handbook on the Socialisation of Scientific and Technological Research, Social Sciences and European Research Capacities, Rome: River Press Group.
- Boylan, J., Dacre, J., Gordon, H. (2019). Addressing women's under-representation in medical leadership. *The Lancet*. 2019; 393(10171): e14.
- Bromme, R. (2000). Beyond one's own perspective. The psychology of cognitive interdisciplinarity. In P. Weingart & N. Stehr (Eds.), *Practising interdisciplinarity* (pp. 115-133). Toronto: Toronto University Press.
- Bubela, T. (2006). Science communication in transition: Genomics hype, public engagement, education and commercialization pressures. Clinical Genetics, 70(5), 445–450. https://doi.org/10.1111/j.1399-0004.2006.00693.x
- Bubela, T., Hagen, G., & Einsiedel, E. (2012). Synthetic biology confronts publics and policy makers: Challenges for communication, regulation and commercialization. Trends in Biotechnology, 30(3), 132– 137. https://doi.org/10.1016/j.tibtech.2011.10.003
- Burchell, K. (2015). Factors affecting public engagement by researchers: literature review, Policy Studies Institute, London, https://wellcome.ac.uk/sites/default/files/wtp060036.pdf (accessed on: 24/07/2019).
- Burns, D., Squires, H., (2011). Embedding public engagement in higher education: Final report of the national action research programme, NCCPE. https://www.publicengagement.ac.uk/sites/default/files/publication/action\_rese arch\_report\_0.pdf (accessed on: 26/07/2019)
- Burns, T.W., O'Connor, D.J., & Stocklmayer, S.M. (2003). Science communication: A contemporary definition. Public Understanding of Science, 12(2), 183–202
- https://doi.org/10.1177/09636625030122004
- Capps, D.K., & Crawford, B.A. (2013). Inquiry-Based Instruction and Teaching About Nature of Science: Are They Happening? Journal of Science Teacher Education, 24.
- Cartwright, N., & Hardie, J. (2012). Evidence-based policy: A practical guide to doing it better. Oxford: Oxford University Press.
- Caulfield, T. (2005). Popular Media, Biotechnology, and the "Cycle of Hype". Houston Journal of Health Law & Policy, 337(2004).
- Chen, H.T. (2012). Evaluation von Programmen und Projekten für eine demokratische Kultur. In R. Strobl, O. Lobermeier, W. Heitmeyer (eds.). Evaluation von Programmen und Projekten für eine demokratische Kultur. Fachmedien Wiesbaden: Springer.
- Clark, B.R. (1998). Creating Entrepreneurial Universities: Organisational Pathways of Transformation. Pergamon, http://blog.ub.ac.id/yogidwiatmoko/files/2012/12/gibb hannon.pdf
- (accessed on: 17/09/2019).

- Clark, H. (1996). Using Language. Cambridge: Cambridge University Press.
- Clarke, L.J., & Kitney, R.I. (2016). Syntheticbiology in the UK-an outline of plans and progress. Synthetic and systems biotechnology, 1(4), (https://www.ncbi.nlm.nih.gov/pmc/articles/PMC5625736/ (accessed on: 18/07/2019).
- Colizzi V., de Oliveira T., Roberts R.J. (2007). Libya should stop denying scientific evidence on HIV. Nature; 448 (7157):992.
- Colizzi,V., et al. (2019). Structural Transformation to Attain Responsible BIOSciences (STARBIOS2): Protocol for a Horizon 2020 Funded European Multicenter Project to Promote Responsible Research and Innovation. JMIR Res Protocl 8(3):e11745, https://www.researchprotocols.org/2019/3/e11745/ (accessed on: 30/09/2019).
- Condit, C. (2001). What is "public opinion" about genetics? Nature, 2(10). https://doi.org/10.1038/35093580
- d'Andrea, L., Marta, F.L., Kahma, N. and Vase, S., (2017). FIT4RRI, Project Report on the Literature Review, Deliverable 1.1 (public), December 31<sup>st</sup>, 2017, https://zenodo.org/record/1434349#.W8iFg3szbcs (accessed on: 30/09/2019).
- d'Andrea, L., & Declich, A. (2005). The sociological nature of science communication. *JCOM*, *4*(2).
- d'Andrea, L., Quaranta, G., & Quinti, G. (2005). Manuale sui processi di socializzazione della ricerca scientifica e tecnologica. CERFE. Rome.
- Dahler-Larsen, P. (2006). Evaluation after Disenchantment? Five Issues Shaping the Role of Evaluation in Society. In Shaw, I.F., Greene, J.C., Mark, M.M. (eds.), *The Sage Handbook of Evaluation*. London: Sage Publications.
- Dahler-Larsen, P. (2012). *The evaluation society.* Stanford: Stanford University Press.
- D'Armiento, J., Witte, S.S., Dutt, K. Wall, M., McAllister, G. (2019). Achieving women's equity in academic medicine: challenging the standards. *The Lancet*. 2019; 393(10171).
- Declich G., d'Andrea, L. (2018), "Triggering Institutional Change Towards Gender Equality In Science. Final Guidelines of the TRIGGER Project", project funded by the European Commission under the FP7 for Research.
- de Oliveira, T., Pybus, O.G., Rambaut, A., Salemi, M., Cassol, S., Ciccozzi, M., Rezza, G., Gattinara, G.C., D'Arrigo, R., Amicosante, M., Perrin, L., Colizzi, V., Perno, C.F. (2006). Benghazi Study Group. Molecular epidemiology: HIV-1 and HCV sequences from Libyan outbreak. Nature. 2006; 444.
- Döring, N., & Bortz, J. (2016). Forschungsmethoden und Evaluation in den Sozial- und Humanwissenschaften. Berlin: Springer.
- Eden, G., Jirotka, M., & Stahl, B. (2013). Responsible research and innovation: Critical reflection into the potential social consequences of ICT. In Research Challenges in Information Science (RCIS), 2013 IEEE Seventh International Conference on. IEEE.
- Elster, D. (2016). Deliverable 5.1 First Interim Report, University of Bremen, Bremen.
- Elster, D., Barendziak, T., Birkholz, J. (2016). Science Education as a Trigger to Attain Responsible Research and Innovation. In Pixel: New Perspectives in

Science Education, Conference Proceedings 2017, Florence/LibreriaUniversitariaEdizioni.

- Elster, D., Barendziak, T., Birkholz, J. (2019). Towards a sustainable and open science. Enhancing responsible research and innovation in the biosciences at the University of Bremen. Bremen: University of Bremen.
- Equality Challenge Unit (2005). Athena SWAN Charter. 7th Floor, Queen's House, 55/56 Lincoln's Inn Fields, London, WC2A 3LJ https://www.ecu.ac.uk/equality-charters/athena-swan/
- ERA CoBioTech (2018). Strategic Agenda a vision for biotechnology in Europe, November 2018, https://www.cobiotech.eu/lw\_resource/datapool/systemfiles/elements/files/7D5 DE99D41EC4DCCE0539A695E869159/current/document/114492\_ERA\_CoBio\_a genda\_final\_high-res.pdf (accessed on 18/07/19)
- ERA CoBioTech (2018). Strategic Agenda a vision for biotechnology in Europe, https://www.cobiotech.eu/lw\_resource/datapool/systemfiles/elements/files/7D5 DE99D41EC4DCCE0539A695E869159/current/document/114492\_ERA\_CoBio\_a genda\_final\_high-res.pdf (accessed on: 18/07/19)
- ESFRI European Strategy Forum on Research Infrastructures (2006), Roadmap for European Research Infrastructure. Report of the Biology and Medical Science. Roadmap Working Group, October 2006, https://ec.europa.eu/research/infrastructures/pdf/esfri/esfri\_roadmap/roadmap \_2006/bms-report-roadmap-wg-2006\_en.pdf (accessed on: 05/09/2019).
- European Commission (2016). SheFigures 2015, Directorate-General for Research and Innovation, Brussels.
- Eze, M.O. (2008). What is African communitarianism? Against consensus as a regulative ideal. South African Journal of Philosophy. 27(4). 10.4314/sajpem.v27i4.31526
- Faria, N.R, Quick, J, Claro I.M, Theze, J, de Jesus, J.G, Giovanetti, M. et al. (2017). Establishment and cryptic transmission of Zika virus in Brazil and the Americas. Nature 546.
- Faria, N.R. et al. (2016). Mobile real-time surveillance of Zika virus in Brazil. *Genome Medicine*, 96.
- Felt, U., Fochler, M., & Sigl, L. (2017). IMAGINE RRI: A Card-based Method for Reflecting Responsibility in Life Science Re-search
- Filardo G, da Graca B, Sass DM, Pollock BD, Smith EB, Martinez MA. Trends and comparison of female first authorship in high impact medical journals : observational study (1994-2014). *BMJ* 2016;352:1–8. doi:10.1136/bmj.i847
- Flipse, S.M., Van der Sanden, M.C., & Osseweijer, P. (2014). Setting up spaces for collaboration in industry between researchers from the natural and social sciences. Science and engineering ethics, 20(1).
- France, B., Gilbert, J.K. (2006). A model of communication about biotechnology. Rotterdam: Sense Publishers in cooperation with The New Zealand Biotechnology Learning Hub.
- Gade, C.B. (2012). What is ubuntu? Different interpretations among South Africans of African descent. South African Journal of Philosophy. 31(3). 10.1080/02580136.2012.10751789

- Gardy, J.L., and Loman N.J. (2018). Towards a genomics-informed, real-time, global pathogen surveillance system. Nature Reviews Genetics (19) (1.
- Gerber, A. (2018). RRI: How to 'mainstream' the 'upstream' engagement. *Journal* of Science Communication, 17(3), C06,
- https://jcom.sissa.it/sites/default/files/documents/JCOM\_1703\_2018\_C06.pdf (accessed on: 21/07/2019)
- Gibb, A. and Hannon, P. (2006). "Towards the entrepreneurial University?, in International Journal of Entrepreneurship Education, v. 4, http://blog.ub.ac.id/yogidwiatmoko/files/2012/12/gibb\_hannon.pdf (accessed on: 30/10/2018)
- Gittelman, M., (2016). The revolution re-visited: Clinical and genetics research paradigms and the productivity paradox in drug discovery. Res. Policy, http://dx.doi.org/10.1016/j.respol.2016.01.007
- GREAT (2013). Annual report on the main trends of SiS, in particular the trends related to RRI, http://www.great-project.eu/deliverables\_files/deliverables05.
- Griffiths, R. (2004). Knowledge production and the research- teaching nexus: The case of the built environment disciplines. *Studies in Higher Education* 29, no. 6.
- Healey, M. (2005). Linking research and teaching: Exploring disciplinary spaces and the role of inquiry- based learning. In *Reshaping the University: New Relationships Between Research, Scholarship and Teaching*, edited by R. Barnett. Maiden head, UK: McGraw- Hill/Open University Press.
- Herschberg, C., Benschop, Y., & van denBrink, M. (2018). Precarious postdocs: A comparative study on recruitment and selection of early-career researchers. Scandinavian Journal of Management, 34(4).
- Herzog, C. (2016). Successful comeback of the single-dose live oral cholera vaccine CVD 103-HgR. Travel medicine and infectious disease, 14(4). doi: 10.1016/j.tmaid.2016.07.003.
- Hill, S.C. et al. (2019). Emergence of the Zika virus Asian lineage in Angola. bioRxiv 520437; doi: https://doi.org/10.1101/520437.
- Jagsi, R., Guancial, E., Worobey, C., Henault, L., Chang, Y., Starr, R., Tarbell, N., Hylek, E. (2006). The 'Gender Gap' in Authorship of Academic Medical Literature – A 35-Year Perspective. N Engl J Med 2006;355.
- Jenkins, A. (2004). A Guide to the Research Evidence on Teaching- Research Relations. York, UK: The Higher Education Academy. Available online: https:// www.heacademy.ac.uk/system/ files/ id383\_guide\_to\_research\_evidence\_ on\_teaching\_research\_relations.pdf (accessed on: 20/06/2019).
- Kalpazidou Schmidt, E., Ovseik, P.V., Henderson, L.R., & Kiparoglou, V. (2019). Understanding the Athena SWAN award scheme for gender equality as a complex social intervention in a complex system: analysis of Silver award Action Plans in a comparative European perspective. *bioRxiv. doi:10.1101/555482.*
- Kalpazidou Schmidt, E. & Cacace, M. (2017). Addressing gender inequality in science: the multifaceted challenge of assessing impact. *Research Evaluation*, vol. 26, no 2.
- Kalpazidou Schmidt, E. & Cacace, M. (2018). Setting up a Dynamic Framework to Activate Gender Equality Structural Transformation in Research Organisations. Science and Public Policy, vol. 59.

- Kalpazidou Schmidt, E. (2009). Evaluation, in Bijker W. E. & A'Andrea (eds.), Handbook on the Socialisation of Scientific and Technological Research, Social Sciences and European Research Capacities, pp. 169-189, Rome: River Press Group.
- Kalpazidou Schmidt, E. (2016). Development of monitoring and assessment tools of structural transformation actions to attain responsible biosciences. STARBIOS2 report.
- Kalpazidou Schmidt, E.K., & Cacace, M. (2018). Setting up a Dynamic Framework to Activate Gender Equality Structural Transformation in Research Organisations. *Science and Public Policy*.
- Kuhlmann, S., Lindner, R., & Randles, S. (2016). Conclusion: making responsibility an institutionalised ambition. In *Navigating Towards Shared Responsibility in Research and Innovation: Approach, Process and Results of the Res-AGorA Project* (pp. 161-166). Fraunhofer ISI.
- Kwiek, M. (2015). "Academic Entrepreneurialism and the Changing Governance in Universities. Evidence from Empirical Studies", in Reihlen, W.M., Frost, J., Hattke, F. (eds.) Multi-level Governance of Universities: The Role of Strategies, Structures, and Controls
- Lawrence, T., Suddaby, R., & Leca, B. (2011). Institutional work: Refocusing institutional studies of organisation. *Journal of Management Inquiry*, 20(1).
- Lederman, N. G., Antink, A., & Bartos, S. (2014). Nature of science, scientific inquiry, and socio-scientific issues arising from genetics: A pathway to developing a scientifically literate citizenry. Science & Education, 23(2).
- Lindlof, T.R. (1995). Qualitative Communication Research Methods. London: Sage.
- Lutz, D.W. (2009). African Ubuntu Philosophy and Global Management. *Journal* of Business Ethics. 84(3), 313-328. 10.1007/s10551-009-0204-z
- Mann, A., & Di Prete, T.A., (2013). Trends in gender segregation in the choice of science and engineering majors. *Social science research*, *42*(6), https://www.ncbi.nlm.nih.gov/pubmed/24090849 (accessed on: 26/07/2019)
- March, J.G., Gherardi, S., & Cimmino, S. (1993). *Decisioni e organizzazioni*. Il Mulino, Bologna.
- Mezzana, D. (2018). Some Societal Factors Impacting on the Potentialities of Electronic Evidence, in M.A. Biasiotti et al. (eds.), Handling and Exchanging Electronic Evidence Across Europe, Law, Governance and Technology Series 39, Springer, https://doi.org/10.1007/978-3-319-74872-6\_14.
- Mezzana, D. (ed.) (2011), Technological responsibility. Guidelines for a shared governance of the processes of socialisation of scientific research and innovation, within an interconnected world, Roma, CNR: www.scienzecittadinanza.org/public/SetDevGuidelines.pdf (accessed on: 08/11/2018)
- Msoroka, M.S. & Amundsen, D. (2018). One size fits not quite all: Universal research ethics with diversity. *Research Ethics*. 14(3), 1-17. 10.1177/1747016117739939.
- Musselin, C. (2007). The Transformation of Academic Work: Facts and Analysis. Research & Occasional Paper Series: CSHE. 4.07. *Center for studies in higher education*;

- Naveca, F.G. et al. (2019). Genomic, epidemiological and digital surveillance of Chikungunya virus in the Brazilian Amazon. *PLoS Negl Trop Dis*. 13, 3-0007065.
- Nowotny, H. (2007). Knowledge Production and its Constraints: epistemic and societal considerations, paper presented at the Gulbenkian Foundation, Lisbon, 2007, http://helga-nowotny.eu/downloads/helga\_nowotny\_b58.pdf (accessed on: 18/07/2019)
- Nowotny, H., Scott, P., Gibbons, M., & Scott, P.B. (2001). *Re-thinking science: Knowledge and the public in an age of uncertainty*. Cambridge: Polity.
- Owen, R., Forsberg, E-M., Shelley-Egan, C. (2019). RRI-Practice Policy Recommendations and Roadmaps, RRI-Practice project report. Deliverable 16.2, https://www.rri-practice.eu/knowledge-repository/recommendations/ (accessed on: 21/07/2019)
- Pan-African Bioethics Initiative (PABIN), (2003). PABIN Third Conference: Good Health Research Practices in Africa. Addis Ababa, Ethiopia.
- Pawson, R. & Tilley, N. (1997). Realistic Evaluation. London: Sage.
- Pierini, M. (2008). Le prix de la liberté: Libye, les coulisses d'une négociation, Actes Sud.
- Pinheiro, R., & Stensaker, B. (2014). Designing the entrepreneurial university: The interpretation of a global idea. Public Organisation Review, 14(4).
- Quaranta, G. (1985). L'era dello sviluppo, Franco Angeli, Milano.
- Quick, J., et al. (2017) Multiplex PCR method for MinION and Illumina sequencing of Zika and other virus genomes directly from clinical samples. Nat Protoc, 2017. 12(6).
- Reale, E., Nedeva, M., Thomas, D., & Primeri, E. (2014). Evaluation through impact: A different viewpoint. *Fteval Journal*, *39*.
- Reydon, T.A., Kampourakis, K., & Patrinos, G.P. (2012). Genetics, genomics and society: the responsibilities of scientists for science communication and education. Personalized Medicine, 9(6). https://doi.org/10.2217/pme.12.69
- Rog, D.J. (2012). When background becomes foreground: Toward Context-Sensitive Evaluation Practice. *New Directions for Evaluation*, 135.
- Rome Declaration on Responsible Research and Innovation in Europe https://ec.europa.eu/research/swafs/pdf/rome\_declaration\_RRI\_final\_21\_Nove mber.pdf (accessed on: 26/07/2019)
- Royal Society (2006). *Survey of factors affecting science communication by scientists and engineers*, the Royal Society,
- https://royalsociety.org/~/media/Royal\_Society\_Content/policy/publications/20 06/111111395.pdf (accessed on: 24/07/2019).
- Ruggiu, D. (2015). Anchoring European governance: Two versions of responsible research and innovation and EU fundamental rights as 'Normative anchor points'. NanoEthics, 9(3).
- Sadler, T.D. (2011). Socio-scientific issues in the classroom. Heidelberg, Springer.
- Sambala, E.Z., Cooper, S. and Manderson, L. (2019). Ubuntu as a Framework for Ethical Decision Making in Africa: Responding to Epidemics. *Ethics & Behavior*. 10.1080/10508422.2019.1583565
- Shendure, J., Ji, H., (2008). Next-generation DNA sequencing. In *Nature Biotechnology*, 26.

- Smith, R.D.J., Scott, D., Kamwendo, Z.T., Calvert, J. (2019). An Agenda for Responsible Research and Innovation in ERA CoBioTech. Swindon, UK: Biotechnology and Biological Sciences Research Council and ERA CoFund on Biotechnology
- https://www.cobiotech.eu/lw\_resource/datapool/systemfiles/elements/files/858 86BE9C7161C71E0539A695E865A64/live/document/ERA\_CoBioTech\_RRI\_Fram ework.pdf (accessed on: 18/07/19)
- Spruit, S.L., Hoople, G.D., & Rolfe, D.A. (2016). Just a cog in the machine? The individual responsibility of researchers in nanotechnology is a duty to collectivize. Science and engineering ethics, 22(3).
- Stephan, P. (2013). How to exploit postdocs. *BioScience*, 63(4).
- Stilgoe, J., Owen, R., & Macnaghten, P. (2013). Developing a framework for responsible innovation. *Research Policy*, 42(9).
- Sutcliffe, H. (2011). A report on Responsible Research and Innovation for the European Commission. Retrieved from http://ec.europa.eu/research/science-society/document\_library/pdf\_06/rri-report-hilary-sutcliffe\_en.pdf (accessed on: 30/9/2019).
- Temoshok, L.R., & Wald, R.L. (2008). Integrating multidimensional HIV prevention programs into healthcare settings. *Psychosomatic medicine*, 70(5). doi: 10.1097/PSY.0b013e31817739b4.
- Thézé, J. et al. (2018). Genomic Epidemiology Reconstructs the Introduction and Spread of ZikaVirus in Central America and Mexico. *Cell Host Microbe*. 23.
- Van Belle, S.B., Marchal, B., Dubourg D. and & Kegels, G. (2010). How to develop a theory-driven evaluation design? Lessons learned from an adolescent sexual and reproductive health programme in West Africa. *BMC Public Health*, 10, 741.
- Van Schomberg, V. & van Schomberg, R. (2013). A Vision of Responsible Research and Innovation. In Owen, R., Heintz, M. & Bessant J. (Eds.), Responsible Innovation (pp. 51-74). London: John Wiley & Sons, Ltd.
- Von Schomberg, R. (2019). Why Responsible Innovation. In The International Handbook on Responsible Innovation. A Global Resource. Von Schomberg, R. and Hankins, J. (Eds.). Cheltenham: Edward Elgar Publishing. Forthcoming.
- Watermeyer, R. (2015). Lost in the 'third space': the impact of public engagement in higher education on academic identity, research practice and career progression, *European Journal of Higher Education*, 5:3, http://www.tandfonline.com/doi/pdf/10.1080/21568235.2015.1044546 (accessed on: 24/07/2019).

# ABOUT THE KINARBIOS2 GUIDELINES

This guideline aims to help readers formalize and trigger structural change aimed at introducing appropriate RRI-related practices to their own organisations. This is not a series of prescriptions, but an itinerary of reflection and self-interpretation addressed to different actors within the biosciences. To support this itinerary of reflection and self-interpretation, the document provides...

- a description of a general RRI Model for research organisations within the biosciences, that is a set of ideas, premises and "principles of action" that define the practice of RRI in bioscience research organisations,
- some practical guidance for designing interventions to promote RRI in research organisations in the Biosciences, putting into practice the RRI Model,
- a set of useful practices in implementing the structural change process,
- and information on particular STARBIOS2 cases and experiences, as well as materials, tools and sources, are also provided in the Appendix and in the Annex.

