







# GHTM | ACTIVITY PLAN 2021 MONITORING







### **GHTM | ACTIVITY PLAN 2021 - MONITORING**

### 1 Introduction and executive summary:

The year 2021 was, for GHTM, another year of struggle against the adversities and constraints posed by the SARS-CoV-2/COVID-19 pandemic. Reinforcing our commitment to the mission of producing knowledge on global health and tropical medicine, developing tools and strengthening health systems through excellence in research, training and cooperation for development, GHTM researchers were at the forefront of science and evidence-based knowledge production, that has greatly contributed to the fight against this pandemic, both nationally and internationally.

GHTM researchers were able to maintain the excellent scientific research that characterizes GHTM, using mechanisms of working at distance and maintaining social distance, as well as reinforcing our international cooperation that allowed us to keep projects in progress, maintain scientific productivity and even see new projects approved for funding, projects for the coming years, which we hope post-pandemic. The contributions to the assessment of the impacts of the pandemic on health systems and on the health of populations, in particular vulnerable populations, both in Portugal and in the CPLP space, were deeply monitored and studied in close association with the assessment and study of the adverse consequences for the control of infectious endemic diseases that are one of the main focuses of our work – tuberculosis, HIV, malaria and other opportunistic infections. We also reinforced our role in research in vector-borne diseases and in the health of migrants and travelers, by strengthening GHTM laboratory infrastructure with a new ACL/BCL-3 Security Insectary and by maintaining applied research on the control of these diseases and on health care delivery. It is also worth noting the permanent presence in the media, internet and social networks of our researchers, producing and sharing credible information and enlightening populations about all the science on which the global fight against COVID-19 was based and on which global health is based today. A reflection of the success of all this work are the various merit and excellence awards given to our researchers and their projects, in 20/21.







Our scientific council (GHTM CC) and organizational framework (Figure 1) maintained its structure and aims with two changes. Two researchers left the Center and their coordination responsibilities at GHTM CC being replaced by new members that were invited by the GHTM Scientific Coordinator, with the approval of the GHTM CC and GHTM Director, to take over these coordination responsibilities – Reynaldo Dietz ended his contract with IHMT NOVA at the end of 2021 and was replaced at the coordination of the Individual Health Care Research Group by Marcelo Urbano Ferreira ( <a href="https://ghtm.ihmt.unl.pt/profiles/marcelo-urbano-ferreira/">https://ghtm.ihmt.unl.pt/profiles/marcelo-urbano-ferreira/</a>) and Luis Lapão resigned his affiliation with the GHTM Research Center and was replaced at the coordination of Public Health Information Cross Cutting Issue by Inês Fronteira ( <a href="https://ghtm.ihmt.unl.pt/profiles/ines-fronteira-2/">https://ghtm.ihmt.unl.pt/profiles/ines-fronteira-2/</a>).

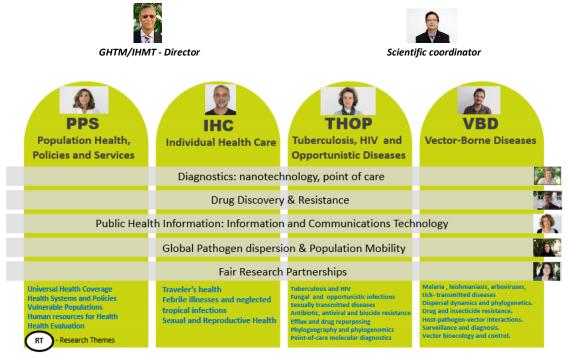


Figure 1 - GHTM / IHMT / NOVA scientific council and organizational framework







In continuation of previous years and despite the limitations, 2021 was a successful year in scientific production and approved projects, with 56 projects in progress and 10 new projects approved. Our 65 researchers (59,70 ETIs) have published 192 international referenced articles, with a weighted citation impact for the scientific area of 22% above the world average. Part of this research was based on the reinforcement of postgraduate training with 11 new doctors in 2021, contributing to the strengthening of our global network of teaching and research in global health and tropical medicine.

- 2 General strategic objectives and 2021/2022 scientific council and organizational framework:
- Promote and develop RESEARCH in the areas of excellence at the center, in the context of the objectives of the research groups (Research Groups -RG) and the priority research lines (Cross-cutting Issues CCIs) of the GHTM / IHMT / NOVA (Figure 1), in line with NOVA's global scientific strategy (emphasizing NOVA Health platform) and IHMT/NOVA:
- Implement in 2021 the strategy defined and approved for the GHTM / IHMT / NOVA for 2019-2023 incorporating the improvement proposals of the FCT Evaluation Panel (2019) and the GHTM Scientific Advisory Board recommendations, ensuring the success of the execution and financing plan approved by FCT for 2019-2023, and implementing specifically the activities and plans approved and compiled at GHTM ACTIVITY PLAN 2021\_03\_02\_FINAL (in attachment to the documentation);
- Promote the **Open Science policy** at GHTM / IHMT / NOVA through the maintenance and constant updating of its activities in the bibliometric repository PURE-NOVA [https://novaresearch.unl.pt/en/organisations/global-health-and-tropical-medicine-ghtm], support for attracting new projects and managing ongoing projects;
- Implement the integrated management GHTM / IHMT / NOVA with the new strategic infrastructures security insectary ACL-3 VIASEF and the Biological Resources Center / biobank Biotropical Resources (BIOTROP) [https://www.ihmt.unl.pt/investigacao/biobanco/];
- Complete the implementation of the general sample management regulation and standard procedures to regulate the data management of the research carried out at IHMT / NOVA;







- Promote the implementation of a **new Associated Laboratory of the Science and Technology Foundation [Associated Laboratory in Translation and Innovation Global Health Towards]**, in consortium with NOVA Medical School's Comprehensive Health Research Center (CHRC) [http://www.nms.unl.pt/main/index.php?option=com\_content&view=article&id=2376:comprehensive-health-research-centre-chrc&catid=2:uncategorised&Itemid=500&lang=pt], the National School of Public Health [https://www.ensp.unl.pt/], FCT / NOVA's LIBPhys [https://www.libphys.fct.unl.pt/] and the University of Évora [https://www.uevora.pt/];
- Framing the research actions of the GHTM / IHMT / NOVA in the Sustainable Development Goals (SDGs) and in the guidelines of the GHTM / IHMT / NOVA;
- Direct and adapt the Central Research Themes of the GHTM / IHMT / NOVA (Figure 1) to the objectives and actions of the European Union and National Horizon Europe Strategies [https://ec.europa.eu/info/horizon-europe\_en] and Portugal 2030 [https://www.portugal.gov.pt/pt/gc21/governo/programa/portugal-2030.aspx] framework and align these themes with the priorities of the Portugal 2030 Strategy and the University NOVA, promoting the sharing of NOVA Saúde initiatives at IHMT / NOVA;
- Align the objectives for Research at the GHTM / IHMT / NOVA with the objectives for the teaching and training of post-graduate human resources at IHMT / NOVA, reinforcing the inclusion and participation of doctoral students (3rd Cycle) of IHMT / NOVA in GHTM / IHMT / NOVA.

In continuation of previous years and despite limitations, 2021 was a successful year in scientific production and approved projects - 56 ongoing projects and 10 new projects approved (3 approved from UE – HORIZON EUROPE - 1 as Coordinator). Our 65 IHMT researchers (59.70 FTEs) (94 GHTM researchers) published 192 international referenced articles, with a weighted citation impact for the scientific area of 1.22% above the world average in 2021 (data consolidation ongoing).

Global output	2018	2019	2020	2021	Evolution
Total of publications	N°	N°	N°	N°	Variation (2018 – 2021)
IHMT	182	175	179	192	<i>&gt;</i>

UO	2016	2017	2018	2019	2020	2021	Variation (2018 – 2021)
	FWCI	FWCI	FWCI	FWCI	FWCI	FWCI	
IHMT	1,33	1,52	1,36	1,11	1.31	1.22 <sup>1</sup>	7

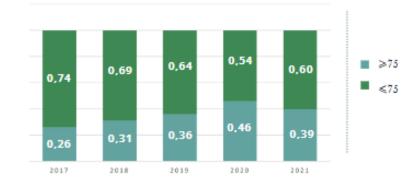
<sup>&</sup>lt;sup>1</sup>data collection for 2021 not consolidated







Figure 2: Percentile 75 distribution of overall publications in the consolidated period (2017-2021)

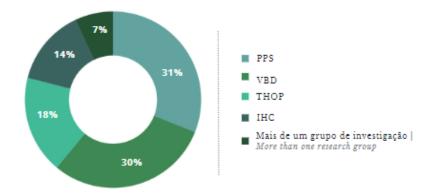


Percentage of publications with international collaboration (2017/2021) 76%

GHTM 2021 publications in SCOPUS TOP 10% 14%

Part of this research was based on the reinforcement of postgraduate training with 11 new doctors in 2021 (9 in 2020), contributing to the strengthening of our global network of teaching and research in global health and tropical medicine.

Figure 3 : Publications distribution by GHTM Research Group in 2021









### **GHTM | ACTIVITY PLAN 2021**

### **MONITORING OF RESEARCH GROUPS ACTIVITIES IN 2021**

- Not done/ Canceled/ No Outcome
- Done/ Concluded/ Outcome
- O- Delayed/ Postponed/ Delayed Outcome







### RG - IHC

main lines of action of the IHC group are related to the transversal areas of the GHTM, Global Pathogen Dispersion and Population Mobility, Drug Discovery & Resistance and Diagnostics and Clinical Trials. The activities planned for 2021 fall within the transversal themes described above:

IHC activities were aligned with Sustainable Development Goals (SDGs) especially #1 – No poverty; #3- Good Health and Well-being, #4-Quality Education and #10 - Reduce inequality within and among countries

ACTIVITIES	PROPOSED/REACHED OUTCOMES FOR 2021	EVIDENCE
Ongoing activities 2020/2021 with CCIs: CCIs - Drug Discovery and Resistance (DDR) and Diagnostics (DG) (Including Clinical Trials)	. 1101 0025/112/101125 00 1001120 1 011 2021	
Therapeutic trials for the treatment of African Human Trypanosomiasis (THA) coordinated by DNDi and EDCTP funded. The 1st is an ongoing Clinical Trial on the "Efficacy of Fexinidazole in the treatment of patients with infection caused by <i>T.brucei rhodesiense</i> in Malawi. The activation of an additional site in Uganda is expected to happen is 2021, which will increase patient enrollment.	(Ongoing) Patient enrollment was completed in October 2021. Last patient-last follow up visit should be completed in October 2022.	Lutje, V,Seixas J. et al. (2021). Chemotherapy for second-stage human African trypanosomiasis: drugs in use. The Cochrane database of systematic reviews, 12(12), CD015374. https://doi.org/10.1002/14651858.CD015374
A second clinical trial with the new oral drug (Acoziborole), in a single dose, for the treatment of THA caused by T.b. gambiense in children and adults (≥15 years), coordinated by DNDi, was recently approved by the EDCTP for funding. This study is expected to start enrolling patients from The Democratic Republic of Congo and Guiné in 2021.	(Ongoing). Due to unexpected difficulties in the development of the pediatric formulation, the clinical trial could only be started in January 2022 in the DRC.	mapo.//doi.olg/10.1002/11001000.00010011
Observational cross-sectional study on visceral leishmaniasis (VL), caused by L. infantum. This corresponds to the Doctoral Thesis of a student in Tropical Medicine, will determine the prevalence of asymptomatic human VL in Portugal.	Thesis topic of a PhD student in Tropical Medicine, scheduled to start in mid-2021	One PhD grant : Rafael Amorim Rocha Supervisors: Carla Maia, Cláudia Conceição, and Luzia Gonçalves. Title: Prevalence of asymptomatic leishmaniasis in Portugal. (SFRH/BD/GHTM/DTM2020)
Surveillance of antimalarial drug response in imported malaria cases in Portugal.	(Ongoing)	Silva-Pinto, A., et al . (2021).  Artemether-lumefantrine treatment failure of uncomplicated Plasmodium falciparum malaria in travelers coming from Angola and Mozambique. International Journal of Infectious Diseases, 110, 151-154.  https://doi.org/10.1016/j.ijid.2021.07.008







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New research Projects:  CCI - Public Health Information (PHI) - COVID-19 vaccination in migrants from Lisbon area: protocol will be developed with the objective of evaluating the knowledge, attitudes, practices and health literacy of migrants with regard to safety, efficacy and confidence in vaccination against COVID 19, which should take place in 2021. The target population will include a sample of migrants from CPLP countries, mainly from the PALOP countries and Brazil, from the Greater Lisbon		Submitted and approved for funding in 2021 and should start in March 2022. Funding: Centro Padre Álvares Correia, Lisboa.
area. –  CCI – Diagnostics (DG) – A cross-sectional study, to be developed at the Hospital de Lubango, Angola, is scheduled for 2021. The objective will be to evaluate the performance of clinical and laboratory diagnostic techniques in the differential diagnosis of SARS CoV-2 infection in an endemic area of malaria, DENV, CHIKV, Salmonella and Leptospirosis and with seasonal incidence of viral and bacterial respiratory infections in winter. It is also intended to develop an optimized diagnostic algorithm, based on the results of the discriminating power of the clinical case definition parameters and the diagnostic tests used.		Study start was postponed mainly due to the low incidence of SARS-CoV-2 in the proposed study site. The study is under revision in order to possibly drop the SARS-CoV-2 component. Inclusion of a second study site in Luanda is under evaluation
Congresses//Events:		
Congresses//Events.		
8th International Conference on <i>Plasmodium vivax</i> Research. Global virtual event. (https://www.vivax2022.org/). IHC members involved.	Event held in 5-8 April, 2022.	
XVII Reunião Nacional de Pesquisa em Malária. Local in-person event, Rio de Janeiro, Brazil, 24-28 April, 2022 (https://www.malaria2020.com.br/). IHC members involved:	Event to be held in 24-28 April, 2022.	
Symposium on Antimalarial Drug Development and Molecular Mechanisms of Resistance. In-person/virtual event jointly organized by IHMT and ICB-USP in São Paulo, Brazil. Organizers: IHC menbers	Event to be held in August, 2022.	
Publications/Training:	At least 5 papers in international journals related with the results and outcomes of the ongoing clinical trials and projects indexed in Scopus.	At least 21 publications in journals indexed in Scopus.
	At least one new doctoral student enrolled in GHTM-IHC group	Rafael Rocha, MD, PhD candidate in the Tropical Medicine program. Ongoing.







### Capacity building:

- Zoom meetings for training local PIs in Lubango, Angola
- Keep updated the PURE-NOVA profiles of IHC members.
- Meeting on human African trypanosomiasis elimination

- Participation in open science initiatives
- Knowledge dissemination initiatives on the ongoing pandemics supporting efforts from the public health authorities, including seminars and interviews for the media.

Participation of several IHC members as scientific/technical reviewers of the digital platform on vaccines and immunization IMUNE.PT, awarded with two prizes of communication/scientific dissemination.

Participation of IHC members in initiatives raising general public comprehension on COVID.

A one-day Workshop-Webinar on 'the sociocultural dimensions and community perspectives on HAT Elimination' was organised by the Institute of Hygiene and Tropical Medicine in Lisbon (Portugal) in collaboration the WHO HAT Elimination Network on 19 July 2021.

Coordination: Philip J. Havik (IHMT); Jorge Seixas (IHMT); Jorge Atouguia (IHMT); José Ramón Franco (WHO); Gerardo Priotto (WHO); Augustin Kadima Ebeja (WHO)

https://www.imune.pt/

JNina and CConceição, among others: Regular presence in the Media – TSF radio, SIC, RTP and TVI Networks, Newspapers and magazines (see 2021 IHMT Report for further details)







### RG - PPS

The main research topics of the PPS group are: Health Policy and Planning; Human Resources for Health; Health determinants of vulnerable populations. The activities planned for 2021 fall into the various cross-cutting areas described above:

PPS activities were aligned with Sustainable Development Goals (SDGs) especially #1 – No poverty; #3- Good Health and Well-being; #4-Quality Education and #10 - Reduce inequality within and among countries.

ACTIVITIES	PROPOSED/REACHED OUTCOMES FOR 2021	EVIDENCES
Ongoing activities 2020/2021 with CCIs:		
CCI - Global Pathogen dispersion and Population Mobility (GPPM) 3 studies related to the adult and child immigrant population:		
at the adult level, a work on acculturation, lifestyles and health of CPLP immigrants in two lberian contexts;	No information	No information
in terms of child health, the cohort study in the Amadora region will be continued through a second survey of data	Data on the second wave of data were collected through phone and face to face interviews and administrative data	Data set completed
the elaboration of the Protocol for a cohort study of children in Cape Verde and / or S. Tomé and Prince.	Cape-Verde: Protocol for the first data collection already approved by Cape-Verde NEC; questionnaires already in redcap; (S. Tomé: not implemented).	Protocol approval
Project on the impact of the pandemic on health inequalities (FCT funding)	Determinants and health needs of migrant children in a pandemic context: a longitudinal study for the Region of Lisbon and the Tagus Valley.	Signed and started 2021 Researcher Fellowships
CCI - Public Health Information (PHI) 2 projects funded by European Funds:  VAX-TRUST - Addressing vaccine hesitancy in Europe –	ragus valley.	Signed and started - 1 Fellowship: Ongoing. Edital already completed 1 Grant Holders /Fellowship.
Financing UE -Horizonte 2020.  BCG vaccine to reduce unplanned absenteeism due to illness of health care workers during the COVID-19 pandemic. A multi-center randomized controlled trial. Financing EU /ECDTP.	Data for some countries already collected.	Signed and started Fellowship1 Grant Holders /Fellowship posdoc
Knowledge dissemination initiatives on the ongoing pandemics supporting efforts from the public health authorities, including seminars and interviews for the media.	Participation of PPS members in initiatives raising general public comprehension on COVID.	Tiago Correia among others: Regular presence in the Media – TSF radio, SIC, RTP and TVI Networks, Newspapers and magazines (see 2021 IHMT Report for detais)







CCI – Fair Research Partnerships (FRP)  2 important partnerships: one between the United Kingdom, Portugal and Mozambique as part of an investigation into patterns of health literacy and demand for men's health care in Mozambique and another between Portugal, Mozambique, Cape Verde and Angola on strengthening the capacity of bioethics committees.		1 postdoc fellowship for 6 months in bioethics
IANDA GUINÉ SAÚDE - Guinea-Bissau Health Reinforcement (Inst Camões, FCG)	Ongoing	
New research Projects:  6 projects (as PI) and 1 approval were submitted to FCT's last call.		
● VAX-TRUST - Addressing vaccine hesitancy in Europe- H2020	Signed in February 2021 and already started	
BCG vaccine to reduce unplanned absenteeism due to illness of health care workers during the COVID-19 pandemic. A multi-center randomized controlled trial - UE/EDCTP	Signed and already started. Most of data already collected	
Non-specific effects of BCG in under-five children"- FCT, EXPL/SAU-EPI/0067/2021	Signed and started	
Health literacy in times of coronavirus and COVID-19: the COVID-HL school survey.	Started and already ended	
Determinants and health needs of migrant children in a pandemic context: a longitudinal study for the Region of Lisbon and the Tagus Valley. PTDC/SAU-SER/4664/2020	Signed in February 2021 and already started	
Impact of COVID-19 on non-EU migrant's health: knowing more for a better intervention-PT/2021/FAMI/699	Signed in October 2021 and already started	
Impact of deforestation for the emergence of new emergent infectious diseases- Marie Curry-H2020	Marie Currie Grant; signed and started in January 2022	
A cohort study of HBV-infected pregnant women in Lubango, Angola: vertical transmission and relationship with maternal, obstetric and newborn complications	Not funded	







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Conferences/events: Organization of 5 National and international seminars / conferences  Organization of a European Conference (EUPHA Health Workforce Research Group 2nd midterm conference 'Health workforce development: disruption, innovation and resilience beyond the COVID-19 crisis'),	Held in June 2021	Link WEB:
Luso-Brazilian Meeting (in Health Assessment and Knowledge Management)  3 seminars on: Health Human Resources, Knowledge Management and Communication Strategies.	Postponed to 2022  No information	
Publications/Training:  PPS group members, including students, will participate, with presentation of communication, in at least 7 Congresses and Conferences:  Publications in journals indexed in Scopus	3rd World Congress on Maternal Fetal and Neonatal Medicine - MARCH 25-27, 2021 - Venice, Italy; International HBV Meeting - September 14-17, 2021 - Toronto, Canada;	Posters: Cá, E., Nielsen, S., Fronteira, I., Benn, C.S., Schaltz-Buchholzer, F. High prevalence of SARS-CoV-2 antibody among health care workers in Guinea- Bissau. Tenth EDCTP Forum, 17-21 de Outubro 2021. Fronteira, I., Schaltz-Buchholzer, F.,
	Tenth EDCTP Forum, 17-21 de Outubro 2021.	Nielsen, S., Araújo, I., Aide, P., Ferrinho, P., Bem, C. Prevalence of observed BCG scar in health care workers from the BCG-COVID-RCT trial. Tenth EDCTP Forum, 17-21 de Outubro 2021.  Amaral, P., Fronteira, I. Profile of
	XXXIX Reunión Anual de la Sociedad Española de Epidemiología y XVI Congresso da Associação Portuguesa de Epidemiología, 7 a 10 de setembro, 2021	traditional chinese medicine users according to ICD-11. XXXIX Reunión Anual de la Sociedad Española de Epidemiología y XVI Congresso da Associação Portuguesa de Epidemiología, 7 a 10 de setembro, 2021
		Amaral, P., Fronteira, I. Profile of traditional chinese medicine users according to differential diagnosis of traditional medicine. XXXIX Reunión







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		Anual de la Sociedad Española de Epidemiología y XVI Congresso da Associação Portuguesa de Epidemiología, 7 a 10 de setembro, 2021
	Congress on Tropical Medicine and International Health: Global challenges in health, migration and equity. Sep 27 - Oct 1, 2021 online, Bergen, Norway	Oral communications: Socioeconomic impact of COVID-19 among immigrants in Lisbon Region, 12th European Congress on Tropical Medicine and International Health: Global challenges in health, migration and equity. Sep 27 - Oct 1, 2021 online, Bergen, Norway
	Profissões da Saúde. NOVAsaúde Health Systems and Policies, 12 Nov 2021. Instituto de Higiene e Medicina Tropical da Universidade NOVA de Lisboa	Professions in Health. NOVASaúde Health Systems and Policies, Instituto de Higiene e Medicina Tropical da Universidade NOVA de Lisboa. 12 Nov 2021
	XXXIX Reunión Anual de la Sociedad Española de Epidemiología y XVI Congresso da Associação Portuguesa de Epidemiología, 7 a 10 de setembro, 2021	How the syndemic perspective can change the way we understand the SARS-CoV-2 pandemic. Roundtable: Syndemics: the logic of multiple causes. XXXIX Annual Meeting of the Sociedad Española de Epidemiología and XVI Congress of the Portuguese Association of Epidemiology, September 2021
	I Encontro da Liga de Tuberculose da Escola de Enfermagem de Ribeirão Preto (online), March 2021	Tuberculosis in the global context: challenges for Portugal in "Tuberculosis in the global context: challenges for Brazil, Portugal,







ALLERY 25.00		
	AORTIC Cancer in Africa Conference, November 2021	Nigeria, Mozambique and Guinea- Bissau" at the 1st Meeting of the Tuberculosis League of the School of Nursing of Ribeirão Preto (online) , March 2021  Epidemiological profile of esophageal cancer in patients followed at Dr. Baptista de Sousa hospital in S.
		Vicente island, Cape-Verde, between 2010 and 2015. AORTIC Cancer in Africa Conference, November 2021
	publications in journals indexed in scopus	65 publications (see 2021 GHTM Research Report report)
	At least two new doctoral students enrolled in PPS-IHC group	3 PhD grants signed Regina Amado; Reka Cane; Iolanda Alves
Capacity building:  organization of an advanced Statistics course for INS-MISAU, Mozambique; ethics course in health research in e-learning for CPLP	3 advanced courses 1 in person 2 e-learning	Advanced Course on Health Statistics (May 2022, INS-MISAU)  Training course for field agents within the ECTP-project
		e-learning Course on Ethics for Health Research (14-18 March 2022) 368 candidates 50 selected participants from Angola, Cape-Verde, Guine-Bissau, Mozambique and S. Tomé e Principe







Participation in open science initiatives

Participation of several PPS members as scientific/technical reviewers of the digital platform on vaccines and immunization IMUNE.PT, awarded with two prizes of communication/scientific dissemination.

https://www.imune.pt/







### RG – THOP

The overarching aim of the THOP Group is the molecular epidemiology, diagnosis, study of drug-resistance mechanisms, drug discovery and control of tuberculosis (TB), HIV and opportunistic diseases on a Global Health perspective in an era of antimicrobial resistance, chronic immunosuppressive diseases and now the COVID pandemic. Despite the many challenges and uncertainty brought by the ongoing pandemic, THOP members achieved and in some cases, even exceeded, most of the goals established for 2021 and initiated new areas of research. Most of the planned activities not accomplished, mainly due to the pandemic restraints, were rescheduled for 2022-2023

THOP activities were aligned with Sustainable Development Goals (SDGs) especially #1 – No poverty, #3- Good Health and Well-being, #4-Quality Education, #6-Clean Water and Sanitation, #10 - Reduce inequality within and among countries and #15 - Life on Land.

ACTIVITIES	PROPOSED/REACHED OUTCOMES FOR 2021	EVIDENCES
CCI - DRUG DISCOVERY & RESISTANCE (DDR)		
Drug design and development against mycobacterial drug resistant infections (FCT, EUCAST and EU-MepAnti Marie Skłodowska-Curie action)	<ul> <li>Establishment of the genetic diversity of candidate loci linked to <i>Mycobacterium tuberculosis</i> resistance to bedaquiline, delamanid and pretomanid.</li> <li>Multicentre testing of the EUCAST broth microdilution reference method for MIC determination on <i>Mycobacterium tuberculosis</i>.</li> <li>Design of new antitubercular isoniazid derivatives with improved reactivity and membrane trafficking abilities.</li> </ul>	Gómez-González et al. (2021). Scientific reports, 11(1), 19431. <a href="https://doi.org/10.1038/s41598-021-98862-4">https://doi.org/10.1038/s41598-021-98862-4</a> <ul> <li>Schön et al. (2021). Clinical microbiology and infection 27(2), 288.e1−288.e4.</li> <li><a href="https://doi.org/10.1016/j.cmi.2020.10.019">https://doi.org/10.1016/j.cmi.2020.10.019</a></li> <li>de Faria, et al. (2021). Biomedicine &amp; pharmacotherapy 144, 112362.</li> <li><a href="https://doi.org/10.1016/j.biopha.2021.112362">https://doi.org/10.1016/j.biopha.2021.112362</a></li> </ul>
Computational drug discovery and drug repurposing for <i>M. tuberculosis</i> and <i>Neisseria</i> (EU-Marie Curie Action)	- Identification of 57 approved drugs with predicitve activity against Neisseria gonorrhoeae	1 poster ECCMID2021; 1 poster Microbiotec21.
Antibiotic and biocide resistance profiling in major human and veterinary staphylococcal pathogens (FEDER/FCT)	<ul> <li>Antibiotic and biocide profiling for &gt;400 staphylococci.</li> <li>Proposal of 2 new ECOFFs for identification of antibiotic resistant staphylococcal populations (FEDER/FCT, in collaboration with the COST Action ENOVAT).</li> </ul>	2 posters in ECCMID2021 and 4 posters in MicroBiotec21, 1 Msc thesis Costa et al. Microb Drug Resist. 2021 27:1555-59, https://doi.org/0.1089/mdr.2020.0402
Evaluation of efflux-mediated resistance in the major veterinary pathogen <i>S. pseudintermedius</i> .	- Characterization of efflux activity for 155 S. pseudintermedius;     - COwt values for EtBr and TPP; contribution of efflux for FQ MICs	<ul> <li>Msc Thesis, Ramos B, Msc Biomedical Sciences Dec 2021;</li> <li>Leal M, Msc Biomedical Sciences (to be defended March 2022).</li> <li>2 posters MicroBiotec2021</li> </ul>
Phylogenomics of multidrug efflux pumps in staphylococci	- Workplan delayed due to the pandemics. Expected to be completed by mid-2023	







Antimicrobial resistance in S. aureus causing bloodstream infections in children	- Workplan delayed due to the pandemics. Expected to be	1 poster MicroBiotec2021
in Mozambique  Resistance to antiretrovirals	completed by mid-2022.  - European multi-cohort study data on effectiveness of integrase strand transfer inhibitor-based regimens in HIV-infected treatment-naive individuals.	Rossetti et al.; EuResist Network, INTEGRATE study group. J Antimicrob Chemother. 2021;76(9):2394-2399. https://doi.org/10.1093/jac/dkab200
Drug repurposing for inhibition of SARS-Cov-2 replication (FCT).	- Project concluded, report submitted to FCT	1 poster MicroBiotec2021     One manuscript submitted to "Molecules".
CCI - DIAGNOSTICS (DG)		
SARS-CoV-2 serological survey analysis.	- Data on genome-wide diversity of Zika virus - New nomenclature proposal	Seabra et al. Manuscript accepted at Virus Evolution 2022 3 ongoing MSc Thesis
Development of point-of-care tools for SARS-Cov-2 diagnosis (FCT)	No information	
Development of point-of-care tools for drug resistance detection on M. tuberculosis	Proposal of Electrochemical Detection of Single-Nucleotide Polymorphism Associated with Rifampicin Resistance in <i>Mycobacterium tuberculosis</i> Using Solid-Phase Primer Elongation with Ferrocene-Linked Redox-Labeled Nucleotides.	Ortiz, M et al. (2021). ACS sensors, 6(12), 4398–4407. https://doi.org/10.1021/acssensors.1c01710
Microbicidal activity of macrophages exposed to pathogenic spirochetes Borrelia garinii and B. lusitaniae	(Ongoing)	- 1 poster MicroBiotec2021
Optimization and application of an indirect immunofluorescence test as an alternative to the microscopic agglutination test (reference test) for the diagnosis of leptospirosis (with VBD)	(Ongoing)	-
Purification and characterization of exosomes in Portuguese patients chronically infected with HDV	Project concluded, manuscript in preparation.	
<ul> <li>Determination of the transcriptomic alterations in HDV-induced hepatocellular carcinoma (FCT)</li> </ul>	(Ongoing)	-
Development of point-of-care tools for immunonanodiagnosis of <i>Pneumocystis</i> pneumonia.		<ul> <li>4 papers/book chapter – to confirm</li> <li>4 conference/scientific meeting communications</li> </ul>
ldentification and characterization of intestinal parasitic infections in Mozambique.	Data on occurrence of intestinal parasites in fresh horticultural products sold in maputo markets and supermarkets, Mozambique	<ul> <li>Salamandane et al. (2021). Microorganisms, 9(9), [1806].</li> <li>https://doi.org/10.3390/microorganisms9091806</li> <li>4 conference/scientific meeting communications</li> </ul>
Study of the gut mycobiome using nanopore sequencing approaches	No information	-
CCI - GLOBAL PATHOGEN DISPERSION AND POPULATION MOBILITY (GPPM)		-
Phylogeography, dispersion of MDR/TB in Brazil	Genomic-based surveillance reveals high ongoing transmission of multi-drug-resistant <i>Mycobacterium tuberculosis</i> in Southern Brazil.	Salvato et al. (2021). International journal of antimicrobial agents, 58(4), 106401. https://doi.org/10.1016/j.ijantimicag.2021.106401







- Establishment of the main clonal lineages causing SSTIs ldentification of prevalent clonal lineages of staphylococcal pathogens -Ferreira et al. Antibiotics 2021, 10(5), 518. doi.org/10.3390/antibiotics10050518 in humans and companion animals for 3 staphylococcal associations with multidrug resistance, biofilm formation and virulence potential species Costa et al. Antibiotics 2021, 10(4), 345. (FEDER/FCT). - Evaluation of biofilm production and virulence potential on doi.org/10.3390/antibiotics10040345 representative strains of these main clonal lineages 2 posters in ECCMID2021, 3 posters in MicroBiotec21, 2 Msc thesis - Sociodemographic, Clinical, and Genomic Characteristics Transmission dynamics of HIV-1 in Europe and in migrant populations Tavares et al. AIDS Res Hum Retroviruses, 2021 of HIV and Tuberculosis Co-Infection Among Migrants in Jan;37(1):34-37. doi: 10.1089/AID.2020.0119. Portugal Pimentel et al. AIDS. 2022 Feb 25. doi: - Patterns of post-migration HIV-1 infection acquisition 10.1097/QAD.0000000000003203 among Portuguese immigrants of different geographical Miranda MNS et al. Pathogens. 2021 Jul 2;10(7):835. doi: 10.3390/pathogens10070835. - Determinants of HIV-1 Late Presentation in Patients Miranda AC et al. AIDS Res Hum Retroviruses. 2021 Followed in Europe. Nov;37(11):846-851. doi: 10.1089/AID.2020.0175. Epub 2021 - Determinants of HIV-1 Late Presentation in a Cohort of Portuguese HIV-1 Patients Two oral presentations and one poster in European HIV and Hepatitis Meeting 2021 One abstract selected for oral presentation in ECCMID2022.. One PhD thesis being finalized. Evolutionary history and drug resistance patterns of HIV-1 in Cabo Verde (Ongoing) One abstract submitted to international conferences; one manuscript in preparation One PhD completed. One manuscript submitted. Epidemic history of HBV and HCV in Portugal (Ongoing) Phylogeography, dispersion of HIV and resistance to antiretrovirals in Portugal, (Ongoing) with a special focus on vulnerable populations (migrants and PALOPs). Horizon 2020, FCT: manuscript submitted Genome-wide diversity of Zika virus: exploring spatio-temporal dynamics to guide a new nomenclature proposal Participation in project DogIPM-Immune precision medicine as a new opportunity (Ongoing) to control canine trypanosomatid diseases **CCI - PUBLIC HEALTH INFORMATION (PHI)** Participation of THOP members in >3 initiatives raising Knowledge dissemination initiatives on the ongoing pandemics supporting efforts J Piedade: Oral communication "Vacinas contra a general public comprehension on COVID. from the public health authorities, including seminars and interviews for the media. COVID-19", webinar "VACINAS COVID 19: VERDADES E MITOS", in the scope of project "IANDA Guiné Saúde" (Feb 2021).







		CCunha among others: Regular presence in the Media – TSF radio, SIC, RTP and TVI Networks, Newspapers and magazines (see 2021 IHMT Report for further details)
CCI - FAIR RESEARCH PARTNERSHIPS (FRP)		
Capacity building initiatives on Covid-19 diagnosis and awareness in Angola and other CPLP countries.	Participation in project IANDA Guiné Saúde	J Piedade: Oral communication "Vacinas contra a COVID-19", webinar "VACINAS COVID 19: VERDADES E MITOS", in the scope of project "IANDA Guiné Saúde" (Feb 2021).
		CCunha. webinars for Benguela and another for Luanda Military Hospital
HIV vertical transmission in Angola (w/ HIC and PPS members)	Evaluation on vertical transmission of sexually transmitted infections to newborns.	Oliveira et al. (2021). J. Infect. Dev. Ctries., 15(10):1547-1550. doi:10.3855/jidc.12731.
TRANSVERSAL:	One strategic infrastructure of GHTM IHMT is coordinated by Two additional GHTM transversal facilities coordinated by The	
Definition of GHTM's Bioinformatics HUB's general SOPs for data storage and management.	GHTM's Bioinformatics HUB's general SOPs for data storage and management prepared (awaiting for final GHTM's CC approval)	GHTM's Bioinformatics HUB's general SOPs for data storage and management (2021 version)  General SOPs for MinION use (2021 version)
Training of GHTM members on MinION and Flow Cytometer	General SOPs for MinION usage defined  Formation provided by THOP members (responsible for GHTM's Bioinformatics Hub) to students and technicians on good practices for MinION usage  Training provided by THOP members on GHTM's Flow	Training on MinION provided for 13 GHTM members  Training provided to several PhD and MSc students (individual practice training)
Collaborate/contribute to GHTM's Bioinformatics HUB's and to GHTM's BRC/Biobank	Cytometer to PhD and MSc students.	Integration of IHMT's mycobacteria collections at GHTM's BRC/Biobank (delayed)
NEW RESEARCH PROJECTS APPROVED / STARTED 2021	THOP implemented 1 new research project (as initially anticipate)	pated)
11 submitted projects (6 more than initially anticipated)		
	Signed in 2021 and already started	







EuCARE - European Cohorts of Patients and Schools to Advance Response to Epidemics (UE   HEurope), 2021-2026  New strategies for drug design and development against mycobacterial drug resistant infections; Multicenter validation of EUCAST broth microdilution reference method for MIC determination on Mycobacterium tuberculosis for Pretomanid (TB Alliance), Delamanid (Otsuka) and Bedaquilne (Jenssen) – AMST-ESCMID-EUCAST	Signed and already started. Most of data already collected	
MARVEL - Minimizing the emergence and dissemination of HIV-1 drug resistance in PALOPs through an evidence-based portable high-throughput sequencing and computational approach.	Signed in 2021 and already started	
Next-generation sequencing to understand the HIV-1 transmission patterns in Angola (ENVOLVE Ciência PALOP Fundação Gulbenkian Ciência)	Signed in 2021 and already started	
"TRAINEXTB" - Targeting Mycobacterial Energy Metabolism to Train the Next Generation of Drug Discovery Researchers MARIE SKŁODOWSKA-CURIE ACTIONS. Doctoral Networks (DN) - Call: HORIZON-MSCA-DN-2021	Submitted. Under evaluation by EU HORIZON	
New tools for point of care diagnosis of MDRTB, Malaria and HIV –Rapid, cost-effective detection of infectious disease and antibiotic resistance -RIDAR (Call LaCaixa 2021).	Submitted. Under evaluation by LA CAIXA	
In silico discovery and biological validation of new drugs against Neisseria gonorrhoeae (CEECI, FCT).	Submitted, not awarded	
Computational drug discovery and drug repurposing for Neisseria gonorrhoeae (FCT/LaCaixa).	Submitted, not awarded	
Target.BIOFILM - Drug repurposing to counteract biofilm and antimicrobial resistance in staphylococci	Submitted, not awarded	
Genome-wide diversity of Zika virus: exploring spatio-temporal dynamics to guide a new nomenclature proposal (Horizon 2020, FCT)	Submitted. Under evaluation by EU HORIZON	
Impact of environmental determinants on respiratory microbiome.	Submitted, not awarded	







Searching for immune signature and miRNAs biomarkers to advance Lyme borreliosis blood based differential diagnosis (with VBD as PI and NOVA Medical School,.	Submitted, not awarded	
<ul> <li>Development of a microRNA/target signature of Cancer Drug Resistance from NOVA Medical School (THOP member as PI in IHMT)</li> </ul>	Submitted, not awarded.	
PARTICIPATION IN CONFERENCES	Participation of THOP members in national/international scien	ntific meetings with presentation of project results
<ul> <li>31st European Congress of Clinical Microbiology &amp; Infectious Diseases 2021</li> <li>Tuberculosis 2021</li> <li>European Meeting in HIV &amp; Hepatitis 2021</li> <li>MicroBiotec 2021</li> <li>15th International Workshop on Opportunistic Protists (IWOP-2021), Czech Republic</li> <li>European AIDS Conference 2021 - EACS 2021</li> </ul>		Participation of 6 THOP members at ECCMID2021 (3 posters) and MicroBiotec21 (9 posters) .  Participation of 3 THOP members on European Meeting in HIV & Hepatitis 2021, 2 oral presentations and 1 poster  1 poster in European AIDS Conference 2021 - EACS 2021 (best poster award)
ORGANIZATION OF SCIENTIFIC EVENTS		
Organization of the National Congress of Microbiology and Biotecnology (MicroBiotec21): several THOP members  GHTM sessions/seminar on: - TB and HIV (nano)diagnosis - Determination of the microbicidal activity of macrophages infected with spirochetes of the Borrelia burgdorferi (s.l.)		
<ul> <li>○ 5<sup>th</sup> GHTM Antimicrobial Resistance Awareness Day (Nov 2021)</li> <li>○ Summer School/Course on antibiotic and biocide resistance in bacteria and OneHealth</li> </ul>	- Postponed for 2022 due to overlap w/ MicroBiotec21 - Postponed for 2022 due to pandemic constraints	
PUBLICATIONS/TRAINING		
Increase the number of doctoral students enrolled in GHTM-THOP group	At least 3 new PhD students enrolled in GHTM-THOP group by the end of 2021 (2 supported by GHTM/FCT PhD grants).	Catarina Morais (GHTM/FCT PhD Grant) Jéssica Antunes (GHTM/FCT PhD Grant) Mª Carolina Ferreira (FCT PhD Grant)
Keep the publication track record	At least 12 research papers and one book chapter published during 2021	> 12 research papers in 2021 1 e-book (Matos & Chiao, ed) 2 book chapters (Matos O; Rodrigues L, et al. Methods Mol Biol. 2021;2314:231-245)







Increase internationalization of GHTM-THOP publications	Increase the number of publications with international partners by 10%, all in open access	
Increase the participation of students in GHTM-THOP publications	At least 7 articles published with PhD/MSc students as co- authors	Partially achieved (<7).
CAPACITY BUILDING		
Participation in open science initiatives	Participation of several THOP members as scientific/technical reviewers of the digital platform on vaccines and immunization IMUNE.PT, awarded with two prizes of communication/scientific dissemination.	https://www.imune.pt/
Initiatives on capacity building activities, using online/other platforms involving MSc or PhDs students, technicians, other	Partially achieved.	







### RG - VBD

The Strategic objective of VBD is the reinforcement of local and global capacity to control vector-borne diseases. Major research competences include molecular, genetic, and eco-epidemiological studies, mechanisms of drug and insecticide resistance, transmission and vector/host-pathogen interactions, host-pathogen microbiome, vector bioecology and control, and development of diagnostic and surveillance/control innovative methods.

Although research activities have been hindered by the COVID-19 pandemic and the year 2021 mostly dedicated to recover from delays in work plans, the productivity of the VBD group exceeded the expectations in almost all expected indicators.

VBD activities were aligned with Sustainable Development Goals (SDGs) especially #1 – No poverty, #2 – Zero Hunger, #3- Good Health and Well-being, #4-Quality Education, #6-Clean Water and Sanitation, #10 - Reduce inequality within and among countries and #15 - Life on Land

AATIVITIEA		EWDENOE
ACTIVITIES  ONCOING PEGFAPOU ACTIVITIES IN 2024 MITHIN COL.	PROPOSED/REACHED OUTCOMES FOR 2021	EVIDENCE
ONGOING RESEARCH ACTIVITIES IN 2021 WITHIN CCIs:	Development of research activities in the framework of	
CCI - Diagnostics (DG)	27 financed projects (10 as proponent institution/investigator	). 
CGI - Diagnostics (DG)		
Xenodiagnosis in domestic cats (Felis catus) naturally infected by Leishmania infantum (FAPESP)	ongoing	Vioti et al. doi: 10.1111/tbed.14216
Optimization of an infection protocol by <i>Borrelia burgdorferi</i> s.l. and human macrophages + study on <i>Leptospira</i> spp – ( <b>WITH THOP</b> )	delayed	MSc thesis (Feb2022)
SUBFAM protocol: Submicroscopic <i>Plasmodium falciparum</i> infections in Guinea-Bissau: a barrier to eliminate malaria? (with Bandim Health Project, Guinea-Bissau) – ( <b>ALSO WITHIN CCI FRP</b> )	ongoing PhD thesis of a local student	
External collaborations beyond vector-borne diseases  Virologic screening of fish specimens captured along the Portuguese coast using a tetraplex qPCR to detect and quantify NoV GI, NoV GII and HAV genomes, and the detection of NoV genomic RNA in a pool of seabass brains (with iBET)	Assessment of the circulation enteric viruses in seawater or fish contamination	PhD thesis (non-IHMT) Filipa-Silva et al doi: 10.3390/microorganisms9061149
CCI – Drug Discovery & Resistance (DDR)		
Drug discovery & resistance		
A multitarget strategy to hit all stages of the Plasmodium life-cycle (FCT)	Delayed Manuscript in prep (da Silva et al Synthesis of antimalarial indoles via one-pot A <sup>3</sup> coupling and domino cyclization.)	Milheiro et al doi: 10.1021/acs.inorgchem.0c01795
AMAZING - AMAZonian snake toxins: CreatING value from bioresources (CIRCNA)	Delayed Manuscript in prep	
A small couple against the big three (FCT)	Surface-active ionic liquids (SAIL) obtained by acid-base combination of primaquine with natural fatty acids are disclosed both as antimalarial drugs and drug delivery systems	Silva et al. doi: 10.3390/ijms21155334  MSc thesis, (March 2022)







Pyrazino (1,2b) quinazoline-3,6-diones derivates, their production and uses thereof (Patent P2020, SAICT)		Long et al. doi: 10.1021/acsmedchemlett.1c00589
Development of nanovectors for the targeted delivery in <i>Anopheles</i> mosquitoes of agents blocking transmission of <i>Plasmodium</i> parasites (UE / Euronanomed III)		Lantero et al. doi: 10.3390/biom10081136
Identification of new antimalarial treatments through a target-centred "drug repositioning" approach (FCT)		Cassiano et al Book chapter Tavella et al. doi: 10.1021/acsinfecdis.0c00454 Lima et al doi: 10.1002/cmdc.202000685
Synthetic peroxides as new potential anti-Leishmania Chemotypes (FCT)	1,2,4-trioxolane and 1,2,4,5-tetraoxane endoperoxides against Old-World Leishmania parasites: in vitro activity and mode of action.  Synthesis, structure and antileishmanial evaluation of endoperoxide-pyrazole hybrids.	Two manuscripts 2022 (Pharmaceuticals / Molecules (under review)
Insecticide discovery & resistance  Evaluation of the effectiveness of the environmental repellent 3-(N-Acetyl-N-Butyl)  Aminopropionic Acid Ethyl Ester (IR3535®), as a tool for malaria control (FBA) – (WITH PPS)	In 2022 Participatory mapping of mosquito breeding sites (May 2022) CAP (June 2022) 1st year trial report (nov 2022)	
Host/Vector – Pathogen interactions and Microbiome  Exploring the tick-host interactome: on the path to vaccine development (FCT)		Pacheco et al. doi: 10.1016/j.heliyon.2021.e06721 Couto et al doi: 10.3390/biomedicines9040363 Paulino et al. doi: 10.3390/pathogens10020167 Sanches et al. doi: 10.3389/fcimb.2020.611113
Immunogens compatible with integrated management strategies in tick control (CYTED)	On going	
The Ixodes ricinus group of ticks in the western Mediterranean region and north Africa: new insights into their population genetics and microbiome fauna (FCT)	On going	
Vaccine for Prevention and Treatment of Trypanosoma cruzi Infection (H2020)	Experimental immunization of dog model (prophylactic setting) and humoral and cellular immune evaluation completed.	
Achieving new frontiers through trypanosomatid exosomes (TEx) (FCT)	On going	
When the host cell is not so cosy anymore? A drop off in energy or an increase in toxicity? (FCT)	Host glycolytic metabolite 2,3-DPG impairs P. falciparum intraerythrocytic developmental cycle in in vitro cultures (inhibition of parasite growth, significantly lower progeny)	Morais et al. doi: 10.3389/fcimb.2022.840968  MSc thesis







Identification of genetic modifiers in children with Sickle Cell Anaemia in an endemic area of Malaria, Angola		Manuscript in prep (Carvalho et al 2,3-DPG and the protective effect of pyruvate kinase deficiency against malaria infection – exploring the role of the red blood cell membrane.)  Santos et al doi: 10.1007/s11033-020-05628-8
Effect of Oral Polio Vaccine on gut and upper respiratory microbiomes of infants –	Ongoing financed by GHTM (UID/Multi/04413/2013, FCT, Portugal).	Oral polio revaccination is associated with changes in gut and upper respiratory microbiomes of infants (npj Biofilms and Microbiomes, under review)  Presentations at scientific meetings (IV International Conference NOVAhealth Chronic Disease and Infection "Microbiota and Health" and Microbiotec – Webconference)
CCI - Fair Research Partnerships (FRP)		
Learning to Investigate by Field Experiment for Southeast Asia Emerging Diseases (Erasmus+)	ended 2021	
UMAP (University of Maryland-Angola-Portugal) Malaria working group development (Presidents' Global Impact Fund) (with INIS Angola, University of Maryland, USA)	ongoing PhD thesis of a local student - Study of P. falciparum demography in southwestern Africa, with a focus in Angola. manuscript in prep (Malaria in Angola, in 2021: recent progress, challenges and future opportunities)	poster presentation at 2021 Annual Meeting of ASTMH (Plasmodium falciparum population structure in southwestern Africa:Initial genome wide sequence data from Angola)
■ IANDA GUINÉ SAÚDE - Guinea-Bissau Health Reinforcement (Inst Camões, FCG)	Ongoing	
CCI - Global Pathogen dispersion and Population Mobility (GPPM)		
A global alliance for Zika virus control and prevention (H2020)	ended 2021	
Review on leishmaniasis in the European Union, the Enlargement countries and the European Neighborhood Policy countries (ECDC)	ended 2021 A questionnaire survey of animal and human health authorities in Europe revealed that: i) leishmaniases are considered emergent diseases in most countries and not notifiable in all	Berriatua et al. doi:10.3201/eid2706.210239







	with autochthonous cases; ii) few countries implement surveillance and control targeting both animal and human infections.	
An operational early WARning system for DENgue and other arboviral diseases in Madeira Island		
<ul> <li>ARBOMONITOR: Dengue, Chikungunya and Zika vector in Europe: cost effective optimization of surveillance and control (FCT)</li> <li>Aedes invasive Mosquitoes (EU / COST)</li> <li>A study on the distribution of the kdr mutations V1016G and F1534C, associated with resistance to pyrethroid insecticides, in European populations of Aedes albopictus.</li> </ul>	High dispersal of these mutations in the European continent, which raises alerts about the need to improve monitoring of insecticide resistance in Europe.  Molecular detection assays revealed widespread superinfection of Aedes albopictus with the endosymbiont Wolbachia	
<ul> <li>Molecular characterization of Afrotropical mosquitoes, potential arbovirus vectors using multiple genetic markers and a combination of sequence analysis techniques</li> </ul>	Use of molecular data as analysis of male/female mosquito genitalia for comparative classification of mosquito vectors of multiple arboviruses  Manuscript in prep	MSc thesis     247 Coxl sequences submitted to GenBank / mounted genitalia deposited in the Insect collection of IHMT
Utilization of multiple genetic approaches based for the analysis of nucleotide and protein sequences of classical insect-specific flaviviruses (cISF): characterization of multiple genetic lineages	1st reconstruction of cISF evolutionary history and spatiotemporal dispersal - dynamic and dependent on human activities.	PhD thesis
Assessment of the role of domestic cats as potential reservoirs of zoonotic leishmaniosis	Evaluation of the role of cats as reservoirs for Leishmania:	PhD thesis Pereira et al doi: 10.1016/j.vetpar.2021.109531
Evaluated the utility of mitochondrial cytochrome oxidase subunit I (COX1) and 16S ribosomal DNA (16S-rDNA) sequence analyses as a complementary/alternative tool to classical taxonomy of hard tick species from Portugal	Assessment of the performance of different genetic markers for molecular classification of ticks	Filipe et al https://doi.org/10.1016/j.vprsr.2021.10 0551
Screening of potential zoonotic pathogens in mosquitoes and other arthropods Portugal (with /FacMedVet, Univ Estremadura, Spain)	Detection of three <i>Gongylonema</i> sp. larvae in two specimens of the dung beetle <i>Geotrupes mutator</i> (Marsham, 1802) from western Spain.	Bravo-Barriga et al. doi: 10.21307/jofnem-2021-050
External collaborations beyond vector-borne diseases  Screening and genetic analysis of Human polyomaviruses (HPyVs) in water samples (waste+environmental) the Lisbon Metropolitan area (with iBET)	Developing a two touch-down PCR multiplex protocol for the characterization of HPyVs in the Lisbon Metropolitan area + completion of a MSc Dissertation with a publication (see below).	MSc thesis Condez et al. doi: 10.3390/pathogens10101309.







CCI - Public Health Information (PHI)		
MosquitoWeb (NOVA Health): European Citizen Science project involved in the NEWSERA network (newsera2020.eu/)	Report of the presence of mosquitoes in Mainland Portugal / Characterization of dispersion, activity of both invasive and native species.	https://newsera2020.eu/labs/citizen- and-society-at-large/
healTh RIsk and social vulnerability to Arboviral Diseases in mainland Portugal (FCT)	Started in 2021	
TRANSVERSAL:	Two strategic infrastructures of GHTM IHMT are coordinated by biobank (BIOTROP) and the high security insectary (VIASEF)	members of the VBD group – the
Biotropical resources - BIOTROP (Biobank / Biological Resources Center)		
Activity of 2021 is detailed in the <u>annual report</u> available for consultation at the GHTM IHMT site	Higher visibility and accessibility of biological collections both to internal and external users.  Establishment of a clearly defined workflow for deposition/requisition of samples  Establishment of SOPs, Forms, Agreements, and Informed Consent  Submission and approval by the Ethical Committee of IHMT (CEIHMT)  Project, workflow, SOPs, posters, and flyers for consciousness of users of the ADMT/Traveler's Consultation etc, approved by CEIHMT	http://ihmtweb.ihmt.unl.pt/download/comunicacao/BIOTROP Annual Report_2021_v21-12-2021.pdf
A <u>catalog</u> has been created for internal and external consultation of biological collections		https://www.ihmt.unl.pt/investigacao/biobanco/
Elaboration and implementation of strategic plan for 2021-2023		http://biotropical.ihmt.unl.pt#  http://ihmtweb.ihmt.unl.pt/Download/BI OTROP/BIOTROP 2021- 2023_STRATEGIC_PLAN_vsf.pdf
Collaboration with the Travel Consultation and Central Laboratory of IHMT – Prospective collections [storage of an aliquot of positive samples for any infectious agent, collected during the diagnostics routine, and donated after informed and voluntary consent of patients in post-travel consultation]	NOT IMPLEMENTED	https://www.ihmt.unl.pt/investigacao/biobanco/ (Annex II of Strategic Plan)







Visibility and Participation in Conferences  ☐ Improvement of BIOTROP section at the IHMT website ☐ For information in English to better reach international researchers, the site of NOVA university is being used ☐ Implementation of a communication plan and presence on the social networks		https://www.ihmt.unl.pt/investigacao/biobanco/ https://impact.webview.pt/2021/03/05/biotrop-biotropical-resources-biobank/https://www.facebook.com/biotropicalresources/ https://www.instagram.com/biotropbiobank/ https://www.ihmt.unl.pt/doencastropicais/videos/
<ul> <li>Biobanking African Pediatric Upper Respiratory and Gut Microbiomes.         Poster presentation at Microbiotec'21 Congress, Lisbon, Portugal.         Biotropical Resources – GHTM/IHMT-NOVA Biobank: Biobanking African Pediatric Upper Respiratory and Gut Microbiomes".         IV International Conference NOVAhealth Chronic Disease and Infection "Microbiota and Health", Lisbon. Portugal.     </li> <li>Outreach activities</li> <li>CIÊNCIA VIVA21 Training Secondary School Students. 19-23 July 2021</li> </ul>		https://microbiotec21.organideia.pt/abs tract-book/  https://ghtm.ihmt.unl.pt/ghtm-for- teens-biobanks-discover-their- importance-in-scientific-research/
INFRASTRUCTURE - VIASEF - In vivo Arthropod Security Facility (FCT)	Conclusion in 2021	
The priority was given to the establishment of SOPs and implementation of promotion and dissemination activities, with the objective of attracting funding and research projects as well as to the  Establishment of protocols/SOPs  Development of protocols for <i>Plasmodium falciparum</i> gametocyte production  Other SOPs delayded  Business plan delayed	Construction and implementation concluded in 2021 Operational implementation delayed Business plan and operational management delayed	Building/Facilities concluded https://www.ihmt.unl.pt/viasef-in-vivo-arthropod-security-facility-infraestrutura-de-alta-seguranca/ Inauguration – June 2021 https://www.ihmt.unl.pt/ihmt-nova-infraestrutura-de-alta-seguranca-viasef-in-vivo-arthropod-security-facility-inaugurada-a-21-de-junho-14h30/







Training  Visibility		MSc projects (1 concluded and 2 ongoing) on new larvae diets for Anopheles, evaluation of insecticide resistance for the Guinea-Bissau malaria control program, Aedes albopictus response to environmental alterations  Promotion video https://www.youtube.com/watch?v=EN IfloQV-GE
NEW RESEARCH PROJECTS APPROVED / START 2021	● VBD implemented 6 new research projects (5 more than init	ially anticipated)
Repel+ New Mosquito Repellent Solutions for Malaria Control (Patent P2020) [CCI - Drug discovery & resistance (DDR)]	Approved 2021 / Start 2022	
West African Network for TB, AIDS, and Malaria - WANETAM 3 (EU/EDCTP) [CCI - Fair research partnerships (FRP)]	Contract signed. Started in 2021	
DogIPM - Immune precision medicine as a new opportunity to control canine trypanosomatid diseases (FCT) [CCI - Global pathogen dispersion and population mobility (GPPM)]	Contract signed. Started in 2021	
Tackling trypanosomiasis cardiomyopathy with microRNAs and tridimensional (3D) models - miT3D (FCT – PeX) [CCI - Global pathogen dispersion and population mobility (GPPM)]	Delayed	
MosqDiet - Rearing Anopheles mosquitoes without blood for malaria research and control (FCT AGAKHAN 541725581 / 2019)	Collaboration beyond CPLP (with Burkina Faso, Tanzania and Mozambique) To evaluate the Bloodless diet (US and EU patent: PCT/IB2019/052967 / WO/2019/198013 / US2021030025A1) in different settings (insectaries/species/	
Blodless Diet	Potential entrepreneurship collaboration with NOVASBE collaboration with PROCHEM, Dieburg – Germany, for diet production	
RESEARCH PROJECTS UNDER EVALUATION OR SUBMITTED IN 2021	23 submitted projects (19 more than initially anticipated)	
<ul> <li>Cited network IBEROAMERILEISH: Cooperacion iberoamericana para combatir la leishmaniosis mediante estrategias eficientes.</li> <li>HUdrepMAL - Hydroxyurea treatment for children with sickle cell anaemia– a good companion against malaria anyway (AKDN/FCT)</li> </ul>	Not funded	







Implementation of a malaria genetic and genomic surveillance pipeline for Angola that leverages COVID19-associated infrastructure development (Bill & Melinda Gates foundation, GCE)		
Genomic epidemiology of silent malaria in Guinea-Bissau (La Caixa)		
INOVEC- A research and InNOvation Partnership for enhancing the surveillance and control		
of mosquito VECtors of emerging arboviruses (HORIZON-MSCA-2021-SE-01-01; Proposal number: 101086257)		
Artificial intelligence approaches to accelerate antimalalarial drug discovery (FCT)		
Estratégias em quimiogênomica computacional e biologia sintética para identificação de novos compostos contra Plasmodium vivax (FAPESP)		
ldentification of the mechanisms of resistance to primaquine and tafenoquine in malaria parasites through experimental evolution, genomics and transgenesis (FCT)		
Purydrazides, a novel class of antimalarial drugs (FCT)		
Diagnostics of Low-density malaria infections: a threat to malaria elimination efforts? (FCT)		
Innovative devices for clean and sustainable mosquito control (FCT)		
Portuguese scientific biological collections of interest for health sciences (FCT)		
Point-of-care diagnostic tests for aquaculture-fish pathogens: contribution of in-silico tools for the characterization of microbial surface proteins (PTDC/ASP-PES/0894/2021).		
Microbiome of Phlebotomus perniciosus, the main vector of leishmaniasis in the western		
Mediterranean and the effect of its microbiota on transmitted human and animal pathogens (PTDC/SAU-PAR/3191/2021).		
Mosquitoes Virome: Metagenomic approach to assess the abundance and diversity of virus in mosquitoes isolated from Portugal continental (PTDC/BIA-VIR/0358/2021).		
PARTICIPATION IN CONFERENCES	Participation in at least 10 national/international scientific me results	etings with presentation of project
IV International Conference NOVAhealth Chronic Disease and Infection "Microbiota and Health", Lisbon		
Congress of Microbiology and Biotechnology "Microbiotec – Webconference", Lisbon		
14th International Symposium on Ticks and Tick-borne Diseases, Freie Universität Berlin		
31st European Congress of Clinical Microbiology & Infectious Diseases		
XXXIIIrd EAVA Congress, Belgium		
Il Conferência científica sobre a COVID-19 (webinar)		
2021 CVBD® WORLD FORUM TOUCHPOINT. online		
28th International Conference of the World Association for the Advancement of Veterinary		
Parasitology		







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<ul> <li>Annual Meeting of American Society of Tropical Medicine and Hygiene. online</li> </ul>		
Reunião Anual Sociedade Portuguesa de Genética Humana. online		
ORGANIZATION OF SCIENTIFIC EVENTS	Organization of at least 3 meeting events.	
Portuguese microBiological Resource Center Network meeting, 7th May 2021 (within Pt-mBRCN)	Season 1 (17 March – 22 September; 11 sessions; all RGs)	
Internal scientific sessions – GHTM sessions (Investing in diagnostics towards better health)  GHTM Spirochetes meeting - participation in GHTM Session "Determination of the microbicidal activity of macrophages infected with spirochetes of the <i>Borrelia burgdorferi</i> sensu lato complex (s.l.) – with <b>THOP</b> )		
5 <sup>th</sup> GHTM Antimicrobial Resistance Awareness Day		
Webinar on Diagnosis of neglected tropical diseases: differential diagnosis of febrile syndromes.		
Webinar on trypanossomatid exosomes		
PUBLICATIONS	50 research papers, 1 book and 8 book chapters were published during 2021 Full list is available at 2021 GHTM RESEARCH REPORT	
TRAINING	VBD group have students of different levels (BSc, MSc, and PhD) developing their thesis plans in the framework of ongoing research activities and projects.	
PhD students	At least	
Concluded - 3		
Ongoing - 3		
MSc students		
Concluded - 3 Ongoing - 2		
BSc students		
Concluded - 1		
Ongoing -		
External collaborations beyond vector-borne diseases		
1 MSc concluded (with iBET)		







CAPACITY BUILDING		
Organization of workshops/courses	Cancelled due COVID restrictions on travel/stay	
Hands-on Workshop on "Gene editing in Trypanosomatids: Modern approaches for		
identification of novel drug targets".		
Hosting of an Erasmus+ training student from Spain on Leishmaniasis.		
Hosting of one Short Term Scientific Mission in the framework of the AIM-COST action on		
"Pyriproxyfen for Aedes albopictus control"		
Hosting of Two Short Term Scientific Missions in the framework of the AIM-COST action on		
"Population genetics of Aedes albopictus in Romania and Czech Republic"		
UCIMI/IHMT course on insecticide resistance surveillance for malaria vectors in São Tomé		
and Principe.		
Workshop on flow cytometry		
Effective Communication Skills – 2 <sup>nd</sup> Ed. (NOVA Doctoral School).		
Participation in training actions		
Monoclonal Antibodies Unit (30 h), Master in Biotechnology, University Eduardo Mondlane,		
Maputo, Mozambique		
Three workshops of the European Consortium NEWSERA with the citizen science initiative		
MosquitoWeb.		
Erasmus+ training on gene editing in Leishmania, Welcome Centre for Integrative		
Parasitology, University of Glasgow, UK, Sept-Oct-2021		
System Biology approach towards the characterization of tick–parasite interactions.		
Conferência/aula integrada no Mestrado em Microbiologia, Universidade de Aveiro. 9th June, 2021		
BioSocial Talks, Research Centre for Anthropology and Health (CIAS), Coimbra University		
7th National Conference on Health and Technology, Luanda, Angola		
Hosting of national/international trainees		
1 Short-Term Scientific Mission Grant, Morphological and phylogenetic study of other species		
of Aedes genus that are detected during the surveillance of Aedes albopictus. Current status in		
Europe, September 2021)		
1 Erasmus + Mobility Agreement Staff Mobility for Training, from Poland, 3 months, Tick and		
parasite epidemiology		
1 Erasmus + Mobility Agreement Staff Mobility for Training, Poland, 2 weeks, qPCR detection		
of tick parasites		
1 Erasmus + Programme (PhD student), Poland, 1 month, Tick and parasite epidemiology		
Highschool "Ciência Viva" short trainings		
<u> </u>		







### **GHTM | ACTIVITY PLAN 2021**

### **MONITORING OF CROSS CUTTING ISSUES ACTIVITIES IN 2021**

- Not done/ Canceled/ No Outcome
- Done/ Concluded/ Outcome
- O- Delayed/ Postponed/ Delayed Outcome







### CCI - DIAGNOSTICS (DG)

**Objective**: to develop novel diagnostic tools for epidemiological assessment and surveillance, to improve precision treatment and to provide affordable, rapid and sensitive diagnostic tools for epidemiological assessment and surveillance, to improve precision treatment and to provide affordable, rapid and sensitive diagnostic tools for epidemiological assessment and surveillance, to improve precision treatment and to provide affordable, rapid and sensitive diagnostic tools for epidemiological assessment and surveillance, to improve precision treatment and to provide affordable, rapid and sensitive diagnostic tools for epidemiological assessment and surveillance, to improve precision treatment and to provide affordable, rapid and sensitive diagnostic tools for epidemiological assessment and surveillance, to improve precision treatment and to provide affordable, rapid and sensitive diagnostic tools for epidemiological assessment and surveillance, to improve precision treatment and to provide affordable, rapid and sensitive diagnostic tools for epidemiological assessment and surveillance, to improve precision treatment and to provide affordable, rapid and sensitive diagnostic tools for epidemiological assessment and surveillance, to improve precision treatment and to provide affordable, rapid and sensitive diagnostic tools for epidemiological assessment and surveillance, to improve precision treatment and to provide affordable, rapid and sensitive diagnostic tools for epidemiological assessment and surveillance, to improve precision treatment and to provide affordable, rapid and sensitive diagnostic tools for epidemiological assessment and surveillance, and to provide affordable, rapid and sensitive diagnostic tools for epidemiological assessment and sensitive diagnosti

Key subjects: Develop and implement novel tools; Molecular diagnostics; Biomarkers; Point-of-care testing and devices; Lab-on-a-chip technologies; Infectious Diseases

ONGOING ACTIVITIES 2020/2021 WITH RGs:	OUTCOMES	EVIDENCES	
In collaboration with the IHC group			
Differential diagnosis of SARS CoV-2 infection in an endemic area of malaria, DENV, CHIKV, Salmonella and Leptospirosis and with seasonal incidence of viral and bacterial respiratory infections in winter	To evaluate the performance of clinical and laboratory diagnostic techniques in the differential diagnosis of SARS CoV-2 infection in an endemic area of malaria, DENV, CHIKV, Salmonella and Leptospirosis and with seasonal incidence of viral and bacterial respiratory infections in winter at the Hospital de Lubango, Angola. To develop an optimized diagnostic algorithm, based on the results of the discriminating power of the clinical case definition parameters and the diagnostic tests used.	Study start was postponed mainly due to the low incidence of SARS CoV 2 in the proposed study site. The study is under revision in order to possibly drop the SARS Cov 2 component. Inclusion of a second study site in Luanda is under evaluation  GHTM session on the 17th of March 2021	
Diagnosis of malaria parasites (Central Laboratory)		Santos-Reis & Nina doi: 10.20344/AMP.15814	
In collaboration with the PPS group			
Network COST- HARMONY-Novel tools for test evaluation and disease prevalence estimation.	Implementation of Bayesian Latent Class Models (BLCMs) to both the identification and control of pathogens in humans and animals Until October 2023	Oliveira et al doi: org/10.3390/math8081258 Paper in prep w/ Carla Maia (VBD)	
In collaboration with the THOP group			
Pneumocystis diagnosis		<ul> <li>Matos &amp; Xiao. doi: 10.3389/fmicb.2021.701879</li> <li>Matos &amp; Xiao doi: 10.3389/978-2-88971-156-7</li> <li>Szydłowicz &amp; Matos doi: 10.1016/j.pt.2021.07.010</li> <li>Szydłowicz et al. https://doi: 10.1093/infdis/jiab209</li> </ul>	







Microbicidal activity of macrophages exposed to Diagnosis of infections by spirochetes (THOP / VBD) GHTM session on the 22<sup>nd</sup> of September 2021 (Project PTDC/SAU-INF/3114/2021 not funded; submitted again 2022) pathogenic spirochetes Borrelia garinii and B. Iusitaniae Optimization and application of an indirect Development of point-of-care tools for drug resistance detection on *M. tuberculosis* Ortiz et al. doi: 10.1021/acssensors.1c01710 immunofluorescence test as an alternative to the Opota et al. doi: 10.3389/fpubh.2021.666187 microscopic agglutination test (reference test) for the Schön et al. doi: 10.1016/j.cmi.2020.10.019 diagnosis of leptospirosis (with VBD) Not implemented Diagnosis of fungal infections ongoing; 1st phase concluded in September 2021 https://ghtm.ihmt.unl.pt/cascais-municipality-sars-Serological survey of SARS Cov2 in Cascais, Portugal cov-2-sero-epidemiological-survey/ GHTM session on the 6<sup>th</sup> of April 2022 Identification and characterization of intestinal parasitic infections in Mozambique. Data on occurrence of intestinal parasites in fresh Salamandane et al. doi: horticultural products sold in maputo markets and 10.3390/microorganisms9091806 supermarkets, Mozambique Purification and characterization of exosomes in Portuguese patients chronically Project concluded, manuscript in preparation. infected with HDV Determination of the transcriptomic alterations in HDV-induced hepatocellular (Delayed/Ongoing) carcinoma (FCT) In collaboration with the VBD group Detection of pathogens in mosquitoes Guarido et al. doi: 10.3390/v13112148 Manoj et al. doi: 10.1111/mve.12524 Animal Health - Detection of pathogens in companion animals Vioti et al. doi: 10.1111/tbed.14216 Pereira & Maia. doi: 10.1016/j.crpvbd.2021.100035 Human Health – Neglected diseases Ortiz eta I. doi: 10.36660/ABC.20201236 Environmental Health (external collaborations beyond vector borne diseases) Filipa-Silva et a. doi: 10.1016/i.aguaculture.2021.736693 **NEW ACTIVITIES 2020/2021 WITH RGS AND INFRASTRUCTURES: New research Projects** Evaluation of the Lolli Method: a randomized cluster New tools for point of care diagnosis of SARS-CoV-2 [with **THOP**] intervention study to evaluate the effectiveness of the use of the pooled saliva test strategy called "Lolli







	Method" within the scope of the EuCare Horizon Europe Research Project (ongoing)	
New tools for point of care diagnosis of MDRTB, Malaria and HIV –Rapid, cost-effective detection of infectious disease and antibiotic resistance -RIDAR [ with <b>THOP</b> ]	submitted LaCaixa 2021	
Diagnostics of Low-density malaria infections: a threat to malaria elimination efforts? (FCT) <b>[VBD]</b>	PhD project ongoing (GHTM PhD grant Ronise Silva, DTSG) PTDC/SAU-EPI/0094/2021 not funded; submission to GHTM PEx2022	
CONGRESSES//EVENTS:		
Organization of conference/workshop on Universal access to health care for the populations - Molecular Testing in Infectious Diseases" Organization of a webinar on Diagnosis of neglected tropical diseases: differential diagnosis of febrile syndromes	Postponed, to be carried out in 2022 or 2023	
PUBLICATIONS/TRAINING:	Please see above (Evidence column)	
Capacity building: Internal scientific sessions – GHTM sessions (Investing in diagnostics towards better health)	Transfer of knowledge and technology between GHTM researchers Season 1 (17 March – 22 September; 11 sessions; all RGs)	
	Season 2 (ongoing; start on the 12 <sup>th</sup> of January, 5 sessions, 4 more; external researchers)	







### CCI - DRUG DISCOVERY & RESISTANCE (DDR)

**Objective:** to identify new therapeutic drugs (including single-encounter radical cure for malaria) and transmission-blocking, and timely detection and characterization of drug and insecticide resistance related to mutations, efflux or other mechanisms on key pathogens and vectors. In addition, tracking the prevalence and spread of drug resistance mutations allows national treatment policies to be adjusted. **Key subjects:** Chemogenomics, Drug Repurposing; hit-to-lead discovery, Pre-clinical and clinical trials, Resistance mechanisms, Tropical Medicine

ONGOING ACTIVITIES 2020/2021 WITH RGs:	OUTCOMES	EVIDENCES
In collaboration with the IHC group		
Two therapeutic trials for the treatment of African Human Trypanosomiasis (THA) coordinated by DNDi and EDCTP funded:		
Clinical Trial on the "Efficacy of Fexinidazole in the treatment of patients with infection caused by T.b. rhodesiense in Malawi and Uganda	Patient enrolment was completed in October 2021. Last patient-last follow up visit should be completed in October 2022.	
Clinical trial with the new oral drug (Acoziborole), in a single dose, for the treatment of THA caused by T.b. gambiense in children (<14 years), coordinated by DNDi, was approved by the EDCTP for funding	This study was expected to start enrolling patients from The Democratic Republic of Congo and Guinea in 2021. Due to unexpected difficulties in the development of the paediatric formulation, the clinical trial could only be started in January 2022 in the DRC.	
<ul> <li>Doctoral Thesis of a student in Tropical Medicine, will determine the prevalence of asymptomatic human VL in Portugal</li> </ul>	Thesis title: Prevalence of asymptomatic leishmaniasis in Portugal. Started in July 2021	
Doctoral Thesis of a student in Tropical Medicine and Global Health, will evaluate the presence of miltefosine resistance genetic markers, in isolates of L. infantum, obtained from dogs with signs and symptoms of visceral leishmaniasis.	Prospective supervisor resigned with GHTM	
In collaboration with the THOP group		
<ul> <li>Drug design and development against mycobacterial drug resistant infections (FCT, EUCAST and EU-MepAnti Marie Skłodowska-Curie action)</li> </ul>		Gómez-González, P. J.,et al. Scientific reports, 11(1), 19431. https://doi.org/10.1038/s41598-021-98862-4  Schön, T., et al. CMI, 27(2), 288.e1–288.e4. https://doi.org/10.1016/j.cmi.2020.10.019







Computational drug discovery and drug repurposing for *M. tuberculosis* and *Neisseria* 1 paper in preparation. MSc thesis concluded in 2022: 2 MSc Theses started in 2021: "Identification of new drugs for N. gonorrhoeae through a (EU-Marie Curie Action) "Drug repositioning to decrease resistance mediated by strategy of in silico repositioning" efflux pumps in Neisseria gonorrhoeae and other Gramnegative bacteria." Posters: "Evaluation of the antimicrobial activity of compounds André Pina, Filomena Pereira, Liliana Rodrigues. that inhibit energy metabolism in Gram-negative Identification of new drugs for *N. gonorrhoeae* using an *in* bacteria." silico repurposing strategy. ECCMID2021 & Microbiotec21. Articles in peer-reviewed journals: Project not funded by FCT QSAR modelling and molecular docking for drug design for *M. tuberculosis* and One outcome from a 2017 FCT project this subject de Faria, C. F et al. Biomedicine & pharmacotherapy Neisseria regarding *M. tuberculosis* 112362. https://doi.org/10.1016/j.biopha.2021.112362 Antibiotic and biocide profiling for >400 staphylococci. Antibiotic and biocide resistance profiling in major human and veterinary staphylococcal 2 posters ECCMID2021, 4 posters MicroBiotec2021 pathogens (FEDER/FCT) ECOFF proposed for 2 antibiotics Articles in peer-reviewed journals: Proposal of new criteria for identification of antibiotic resistant staphylococcal Costa et al. Microb Drug Resist. 2021 27:1555-59 populations (FEDER/FCT, in collaboration with the COST Action ENOVAT). doi: 10.1089/mdr.2020.0402. Workplan delayed due to the pandemics. Expected to be Evaluation of efflux-mediated resistance in the major veterinary pathogen S. 2 Msc Thesis, Ramos B, Msc Biomedical Sciences completed by end 2022. pseudintermedius. Dec 2021; Leal M, Msc Biomedical Sciences (to be defended March 2022). 2 posters MicroBiotec2021 Workplan delayed due to the pandemics. Expected to be Antimicrobial resistance in *S. aureus* causing bloodstream infections in children in 1 poster MicroBiotec2021 completed by mid 2022. No information Antifungal activity of novel chemical compounds against Candida albicans and other pathogenic yeasts 2 papers integrase inhibitors drug resistance with THOP Minimal information provided Resistance to antiretrovirals Project completed in 2021 - Delayed - 1 article submitted Drug repurposing for inhibition of SARS-Cov-2 replication (FCT). Bento-Cunha, et al. Submitted to Molecules in march 2022: 1 poster MicroBiotec2021 "Drug repurposing for COVID-19: a review and a novel strategy to identify new targets and potential drug candidates." 1 article in preparation.







In collaboration with the <u>VBD group</u>		
Identification of new antimalarial treatments through a target-centred "drug epositioning" approach (FCT)	FCT project ongoing – To end in 2022	Targeting malaria protein kinases. Adv Protein Che Struct Biol. 2021; 124:225-274.  Violacein-Induced Chaperone System Collapse Underlin Multistage Antiplasmodial Activity. ACS Infect Dis. 202 Apr 9;7(4):759-776  Artificial Intelligence Applied to the Rap Identification of New Antimalarial Candidates with Dus Stage Activity. ChemMedChem. 2021 Apr 8;16(7):109 1103  Computational Chemogenomics Drug Repositionin Strategy Enables the Discovery of Epirubicin as a Ne Repurposed Hit for Plasmodium falciparum and P. viva Antimicrob Agents Chemother. 2020 Au 20;64(9):e02041-19  Deep Learning-driven research for drug discovery: Tackling Malaria. PLoS Comput Biol. 2020 Feb 18;16(2):e1007025
Exploring the tick-host interactome: on the path to vaccine development (FCT)	FCT project ongoing – To end in 2022	Comparative analysis of Rhipicephalus tick salivary gland and cement elementome. Heliyon. Volume 7, Issue 4, April 2021, e06721  Probing the Rhipicephalus bursa Sialomes in Potential  Anti-Tick Vaccine Candidates: A Reverse Vaccinology Approach. Biomedicines 2021, 9, 363  Characterization of the Rhipicephalus (Boophilus microplu Sialotranscriptome Profile in Response of Theileria equi Infection. Pathogens 2021, 10, 167.  Comparative Proteomic Analysis of Rhipicephalus sanguineus sensu lato (Acari: Ixodidae) Tropical ar Temperate Lineages: Uncovering Differences Durin Ehrlichia canis Infection. Front. Cell. Infect. Microbiol., 2 January 2021  Oral presentations:  Velez R, et al. 14th International Symposium on Tick and Tick-borne Diseases, 24-26 Março 2021. Organizaci pela Freie Universität Berlin. Digital.







		Moreira L, et al. Tick-borne pathogens from wild lxodes ricinus and an insight of its microbiota from Portugal mainland. MicroBiotec, Dezembro, 2021
■ Vaccine for Prevention and Treatment of <i>Trypanosoma cruzi</i> Infection (H2020)	Experimental immunization of dog model (prophylactic setting) and humoral and cellular immune evaluation completed.	NA
Synthetic peroxides as new potential anti-Leishmania Chemotypes (FCT)	FCT project ongoing – To end in 2022	Manuscript submitted to Pharmaceuticals (under review)







### CCI – PUBLIC HEALTH INFORMATION (PHI)

**Objective**: To support public health decision making through the production of information and political decision-making tools, namely in the area of epidemiological surveillance and early detection systems.

Key-subjects: epidemiologic intelligence & surveillance; knowledge translation; health literacy; health communication - Knowledge translation through effective communication and promotion of health literacy [public health intelligence center].

ONGOING ACTIVITIES 2020/2021 WITH RGS:	OUTCOMES	EVIDENCES
In collaboration with the THOP group		
Oscreening for infectious diseases in mobile populations and travelers, with monitoring and risks of introducing pathogens that cause STDs and diarrhea of travelers from Lusophone countries and their antibiotic resistance profile.	Number of tracing/epidemiological surveys performed	
BCG vaccine to reduce unplanned absenteeism due to illness of health care workers during the COVID-19 pandemic. A multi-center randomized controlled trial.	Number of epidemiological surveillance alerts / notifications According to the deliverables contracted with the financing agencies	
Participation of members in at least 3 initiatives raising general public comprehension on COVID	Public information and awareness related to COVID-19 Pandemics	THOP, IHC and PPS members, among others: Regular presence in the Media – TSF radio, SIC, RTP and TVI Networks, Newspapers and magazines (see 2021 IHMT Report for further details). Several GHTM member participated in media coverage of COVID-19 pandemics Number of scientific publications on COVID in Q1 journals Report for 2021
In collaboration with the PPS group,		
Initiatives supporting efforts from the public health authorities and CPLP related to the ongoing pandemics.	VAX-TRUST - Addressing vaccine hesitancy in Europe BCG vaccine to reduce unplanned absenteeism due to illness of health care workers during the COVID-19 pandemic. A multi- center randomized controlled trial.	At least one partnership protocol for analysis and strengthening of information systems established with a CPLP health institution by December 31 ACSS from PT.
Oreate an Intelligence center until December 31, 2021		







Develop and strengthen the training / capacity building of human resources in the area of public health information systems, with a particular focus on those related to epidemiological surveillance and early intervention.	At least one training course in the field of clinical trial management	80% of GHTM members with GCP certificate
Create a GHTM data repository that allows to enhance the ability to answer public health issues through Big Data Analysis and artificial intelligence (AI) methods	Creation of the repository until December 31 As been mentioned but not discussed in the SC  Establishment of partnerships with other academic institutions of the CPLP and the EU with a view to training in the areas of Big Data Analysis and IA  Some potential partners have been identified	Conducting at least one training in the field of epidemiological surveillance for members of the GHTM and IHMT NOVA community  Course on Fundamentals of epidemics (BCG_RCT). Foresaw for May 2022 (BCG_RCT)
In collaboration with the VBD group		
Initiatives supporting efforts from the public health authorities and CPLP countries related to the new and emerging epidemics and future pandemics.		MosquitoWeb (NOVA Health): in 2021, this initiative will evolve into an integrated Iberian network of citizen science for mosquito surveillance, in collaboration with INSA and Mosquito Alert (Spain)  HealTh RIsk and social vulnerability to Arboviral Diseases in mainland Portugal
In collaboration with the IHC group,		







### CCI - GLOBAL PATHOGEN DISPERSION AND POPULATION MOBILITY (GPPM)

Objective: addresses global dispersion of infectious diseases using transdisciplinary and integrated approaches in collaboration with partners in endemic countries. Our aim is to understand the factors that underpin the spread or containment of infectious diseases globally and locally, particularly environmental changes and population mobility, in control or elimination contexts.

Key subjects: Chemogenomics; Drug Repurposing; Hit-to-lead discovery; Pre-clinical and clinical trials; Resistance mechanisms; Tropical Medicine

ONGOING ACTIVITIES 2020/2021 WITH RGS:	OUTCOMES	EVIDENCES
In collaboration with the THOP group		
At least 8 papers published in international journals on the following subjects:  Serological survey of SARS Cov2 in Cascais, Portugal Evolutionary history and drug resistance patterns of HIV-1 in Cabo Verde Evolutionary history and evolution of transmitted and acquired HIV-1 drug resistance in Cabo Verde Transmission dynamics of HIV-1 in migrants Determinants of HIV-1 late presentation in Europe Integrase inhibitors drug resistance Epidemic history of HBV in Portugal Epidemic history of HCV in Portugal Unraveling mechanisms of biocide resistance in bacteria. Biocide susceptibility profiling and determination of biocide resistance determinants in relevant pathogens, with emphasis on staphylococci; Characterization of biocide stimulons in bacteria Molecular characterization of staphylococci causing skin and soft tissue infections (SSTIs) in animals and humans - a OneHealth approach	More than 8 papers published overall ins the subjects proposed and in other subjects related with global pathogen dispersion and mobility of populations	1 paper transmission dynamics of HIV in migrants 3 papers HIV-1 late presentation 2 papers integrase inhibitors drug resistance 1 paper spatio-temporal dynamics insect specific flaviviruses 1 paper impact of Ebola epidemic in West Africa 1 paper P.falciparum R in Cape Verde 1 paper MDR TB transmission in Brazil 1 paper DR S. aureus At least 2 papers under revision
Submission of at least 2 projects for funding  At least 3 master theses: At least 1 Ph thesis	More than 2 projects approved	EUCARE: awarded ENVOLVE: awarded MARVEL: awarded Africa Research Excellence Fund - awarded Research, Development Fellowship 2021/22: awarded ORIGHAMI: Transmission dynamics of HIV-1 in migrants Europe (financed by ANRS) (  Jeremias Chone: MSc Margarida Marques: MSc Jacqueline da Graça: VIH em imigrantes - MSc







Organization of at least 1 training event		Rute Marcelino - PhD
In collaboration with the PPS group,  Subjects related to mobility of populations and migrants and on the determinants of emergence of pathogens, particularly we will address the problems of deforestation and vaccine hesitancy:  Impact of deforestation for the emergence of new emergent infectious diseases  Acculturation, lifestyle and health of migrants in different CPLP contexts  Addressing vaccine hesitancy in Europe	Two EU projects approved in 2021 and already started  One paper in Preprints:	Implementation of a children cohort in the archipelagos of Cape Verde and São Tomé e Príncipe (PPS) - EDCTP  Mertens, T.; Abecasis, A. Time to See the Forest for the Trees: Protecting Forests Could Prevent Future Pandemics and Help Preserve a Common Planetary Future. Preprints 2020, 2020060316 (https://www.preprints.org/manuscript/202006.0316/v1  VAX-TRUST - Addressing vaccine hesitancy in Europe– HORIZON EUROPE
In collaboration with the VBD group  We will investigate determinants of pathogen dispersion in different CPLP contexts, through three independent projects:  Modelling the risk of introduction of Aedes invasive species and the establishment of arboviral vector-borne infections in Europe  Epidemiology of leishmaniasis as a reemerging disease in Europe  Malaria in PALOP countries	One project approved in 2021 and already started One EU project finalized	Phylogenomics of Malaria parasites and socio-cultural determinants of malaria dispersal in PALOP countries (VBD)  MosquitoWeb Zikalliance: a global alliance for Zika virus control and prevention
In collaboration with the IHC group,		p.overium.
we will continue the previously implemented projects with the largest Travelers clinic in Portugal, established at IHMT		Tracking infectious diseases in mobile populations and travelers, with monitoring and risks of introducing pathogens that cause STDs and diarrhea of travelers from Lusophone countries and their antibiotic resistance profile (IHC).  Molecular markers of resistance in parasitic populations collected in malaria endemic areas (VBD+IHC).







### CCI - FAIR RESEARCH PARTNERSHIPS (FRP)

**Objective:** To actively promote fair and equitable partnerships - with the assumption that we can contribute to increased productivity if: the definition of the research agenda results from joint work; clarification of responsibilities and promotion of mutual learning ("reverse innovation"); there is data sharing and research networks.

**Key subjects**: institutional practices to promote research integrity; knowledge co-production strategies; reverse innovation;

ONGOING ACTIVITIES 2020/2021 WITH RGS:	OUTCOMES	EVIDENCES
In collaboration with the THOP group		
In collaboration with the VBD and THOP group		
Learning to Investigate by Field Experiment for Southeast Asia Emerging		
Diseases (Erasmus+)		
In collaboration with the PPS group		
2 important partnerships:		
- between the United Kingdom, Portugal and Mozambique as part of an		
investigation into health literacy standards and men's health care search in		
Mozambique		
- between Portugal, Mozambique, Cape Verde and Angola on strengthening the capacity of bioethics committees.		
capacity of bioetifics confinitiees.		
In collaboration with the IHC group		
In collaboration with all RG group		
Monitoring and support to the first PhD program in Biomedical Sciences in	Analysis of ongoing partnerships (namely, PhD Course in	——————————————————————————————————————
Angola – Faculty of Medicine Universidade Agostinho Neto (in collaboration with	Biomedical Sciences Faculty of Medicine UAN Angola (+NOVA) - results until 31 December 2021	Thesis: "Analysis of the mechanisms for implementing the
Universidade NOVA de Lisboa).	- results until 31 December 2021	PhD program in Biomedical Sciences in Angola, in
		partnership with IHMT-NOVA", Carolina Valério, Instituto de Higiene e Medicina Tropical, UNL. (supervisor António
		Carvalho, co-supervisor Isabel Craveiro).
		Carvano, de Supervisor isaber oravenoj.
		2021 - Valério C, Carvalho A & Craveiro I (2021).
		Capacity building and north/south partnerships in health:
		the example of the implementation of the doctoral
		program in biomedical sciences in Angola. Annals of the
		Institute of Hygiene and Tropical Medicine - v. 20:
		Supplement no 1 - Perspectives on medical education in Angola.
		Aligola.







New activities 2020/2021 with RGs and infrastructures:  Or Develop and strengthen institutional capacity building in the area of fair and equitable partnerships		Guidelines for standardization of protocols on Fair and Equitable Partnerships (e.g. co-authorship rules, data sharing) - until 31 December (Ongoing)
Capacity building: Organization of training actions.  Workshop on knowledge management  Workshop on communication strategies	Organization of at least 2 training events	PhD Tropical Diseases and Global Health, UC Rotação Temática – Fair Research Partnerships (20 students enrolled; final output: 4 papers on the subject)  Scientific Writing Workshop by Joana Pais Zózimo, PhD   Lancaster University
Congresses/Events: GHTM sessions	Organization of at least 2 meeting events	Extra GHTM Session on 15-06-21 "Developing an Individual-Based Social Practice Approach to International Partnerships", by the researcher Joana Pais Zózimo, PhD   Lancaster University







### STRATEGIC GHTM / IHMT / NOVA INFRASTRUCTURES



### BIOLOGICAL RESOURCE CENTER (BRC) / BIOBANK

Nowadays, BRC and biobanks are central structures for innovation on global health and advance of precision health care. Through biobanks, a variety of biological specimens may be conserved under exceptional quality conditions, anticipating collection needs for prospective applications, mainly related with development or application of new biomarkers, new therapies, or diagnostic methods. The BRC/biobank of GHTM/IHMT – Biotropical Resources. BIOTROP – was created in 2016 to improve research in tropical medicine and infectious diseases, allowing the supply of high-quality biological resources to researchers.

After a long period dealing with structural (facilities, equipment), operational (governance, elaboration of standard procedures, SOP, and forms) and legal issues (protection of individual data, and conservation of biological resources), BIOTROP is now ready to receive, preserve, and distribute biological samples strictly following the ethical and legal issues, under the Nagoya protocol and the international best practice guidelines. Its data management good practices and confidentiality were recognized by the Data Privacy Impact Assessment (DPIA) evaluation performed by NOVA Data Protection Officer (DPO) (2019). Very recently, in 2019, BIOTROP integrated the Portuguese Roadmap of Research Infrastructures, as member of two networks - Portuguese microBiological Resources Center Network IPt-mBRCN / MIRRI-PT - Portuguese microBiological Resources Center Network [https://www.mbrcn.pt/] / Microbial Resource Research Infrastructure—Portugal [https://www.mirri.org/] and National Biobanks Infrastructure (Biobanco.pt, http://biobanco.pt).

BIOTROP has now around 21,000 samples of microorganisms (or derived products) associated to both human and animal health, vectors, and reservoirs. These collections were donated by the principal investigators of both recent and past research projects. A web catalogue is being elaborated to facilitate the exchange of resources between investigators. To obtain prospective collections of infectious agents of tropical diseases from endemic areas, positive samples collected during the routine diagnostics procedures at the central IHMT's laboratory, from patients coming to ADMT's Travel Medicine Appointment, will be stored. The procedures and flowchart of collection and storage of this material are currently under appreciation of IHMT's Ethical Commission and will be implemented during the next year.

In addition to depositing and distributing samples, BIOTROP has given support to researchers and students to elaborate SOPs of collection and processing of isolates and further services, such as pathogen molecular typing or phylogenetic analysis will be put in practice.

Active participation in dissemination activities will continue as well as seek of national and international collaborations in order to gain visibility and gathering funds.

A Strategic Plan for 2021-2023 detailing all these activities was prepared to be implemented at the beginning of 2021.

https://www.ihmt.unl.pt/investigacao/biobanco/

http://ihmtweb.ihmt.unl.pt/download/comunicacao/BIOTROP Annual Report 2021 v21-12-2021.pdf

http://ihmtweb.ihmt.unl.pt/Download/BIOTROP/BIOTROP 2021-2023 STRATEGIC PLAN vsf.pdf

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### BIOINFORMATICS HUB

The GHTM/IHMT Bioinformatics Hub represents a significant opportunity to improve bioinformatics skills and support research for the entire GHTM scientific community.

The high processing and data storage capacity of this infrastructure has been essential to follow the intense speed, complexity and volume of data that have been produced in light of recent technological advances, especially in the area of genome-wide sequencing.

This infrastructure intends to act at four levels in 2021: sequencing, data management, support for research projects and enhancement of collaborative networks. More specifically:

### I. SEQUENCING:

Nanopore sequencing - MinION

Several GHTM researchers expressed their interest to carry out high-throughput sequencing in-house, as well as in fieldwork situations (refer to the Report on the analysis of the questionnaire for the purchase of MinION equipment and technology, from "Oxford Nanopore Technologies"). After evaluation of the available options, the portable equipment for nanopore sequencing MinION Mk1C, from "Oxford Nanopore Technologies", was purchased in 2020. The management of MinION usage, namely concerning the schedule and storage space verification, will be performed by the coordinator of the Bioinformatics Hub, with the support of the other members of the Hub. Action proposals:

- Implementation of the MinION usage regulation;
- Initial support in the use of the MinION equipment to ensure good application practices:
- Management of usage and timing;
- Reassessment of data storage needs

### II. DATA MANAGEMENT:

Research data produced at GHTM/IHMT has been managed individually by each research group, mainly resorting to ad hoc storage and sharing options.

### Action proposals:

- To define a general standard operating procedure for data storage and management;
- To develop data management plans, as well as integration in data analysis systems, for particular GHTM/IHMT projects that require it;
- To regularly reassess the needs for computer infrastructure or access to remote platforms.

### III. SUPPORTING RESEARCH PROJECTS:

Bioinformatics support in the areas of genomics and transcriptomics was given, between 2019 and 2020, to 6 projects on HIV, HPV, Hepatitis, antibiotic resistance, Zika and other arboviruses.

### Action proposals:

- To continue to support the development of projects and the analysis of genomics and transcriptomics data within the scope of ongoing research activities at GHTM;
  - To manage all the bioinformatics equipment usage, assuring access and good practices, and proposing new acquisitions of computer material according to the research needs.







### IV. COLLABORATIVE NETWORKS:

IHMT is part of collaborative networks within the scope of bioinformatics applied to health. The reinforcement of the existing bioinformatics infrastructure will allow to expand the networks and look for new collaborations and funding.

### Action proposals:

- To continue to boost interaction and collaboration with national and international infrastructures and networks, namely, maintaining the participation in the PT-OPENSCREEN Infrastructure, in the REDCap (Research Electronic Data Capture) consortium, in the PrimaryCare@Covid19 and Lab-BioBanco digital platforms and in networks WANETAM-3 (West African Network for TB, AIDS, Malaria) and TB WGseq & Bioinformatic network (London School of Hygiene and Tropical Medicine);
- To strengthen partnerships and collaborative networks and prepare applications for funding;
- To develop exchange programs with CPLP partners.









The "In Vivo Arthropod Security Facility (VIASEF)" is part of the Roteiro Nacional de Infraestruturas de Investigação since 2014. This infrastructure, hosted at the Instituto de Higiene e Medicina Tropical/NOVA, is devoted to the study of arthropods and the pathogens they transmit. VIASEF is specifically adapted to rear and maintain invasive and non-invasive arthropod vector species, transgenic strains and to perform transmission studies of pathogens of human and animal diseases that need to be handled in BSL-3 laboratories.

Access to VIASEF is universal to the scientific and industry communities. This research facility with ACL-2 and 3 insectaries and laboratories (including a BSL-3 lab) provides access to a wide variety of services. Some of these services are standardized offers that can be purchased for either research or commercial purposes. There is also the possibility of submitting proposals for the development of specific studies. Insectaries and laboratories can be rent, with or without technical support.

This ACL-3 facility offers national and international researchers, new opportunities for the development of scientific activities focusing vector-borne diseases such as malaria, dengue and West Nile fevers, leishmaniasis, Chagas or Lyme disease. Many of these infections are neglected and/or (re)emerging diseases of increasing global health concern. This is of particular relevance in the present context of climate, environmental and social changes which tend to expand the geographic ranges and epidemiological patterns of vector-borne diseases. VIASEF, by promoting fundamental and translational research, as well as the development of new products and services addressing these issues, can significantly contribute to the improvement of health, social and economic status of populations. Furthermore, by fostering the training of researchers, health specialists and human resources for industry, VIASEF is also promoting the creation of new jobs and highly differentiated human resources. In this context, the outcomes of the activity of VIASEF will impact the following United Nations Sustainable Development Goals (SDGs):

- SDG 1 (no poverty), as many vector-borne diseases affect primarily the poorest fractions of both urban and rural human populations.
- SDG 2 (zero hunger), as vector-borne diseases of veterinary importance such as animal trypanosomiasis or tick-transmitted protozoans have major impacts in livestock.
- SDG 3 (good health and well-being), due to the enormous impact that vector-borne diseases such as malaria or Aedes borne arboviruses have in human populations worldwide.
- SDG 4 (quality education), being embedded in a University, VIASEF will accommodate students for experimental training in the scope of their (post-)graduate work programs.
- SGD 8 (decent work and economic growth), as vector-borne disease of both medical and veterinary importance as major causes of economic loss, especially in low-income countries.
- SDG 9 (industry, innovation and infrastructure), due to its capacity for testing new products and solutions for vector-borne diseases control.
- SDG 10 (reduced inequities), as some of the major vector-borne diseases still affect primarily tropical and subtropical regions of the globe, as well as low-income countries.
- SDG 13 (climate action), by providing insectary-based controlled environments for experimental studies on the impact of climate change in transmission of vector-borne pathogens.
- SDG 17 (partnerships for the goals), due to the potential of this infrastructure to promote synergies between academy and industry, at both national and international levels.

The activity of VIASEF also aligns with two of the five priorities for Health, Clinical and Translational Research, of the FCT Research and Innovation Agenda for Health 2020-2030, namely:

- Pharmacology, Medicines and Advanced Therapies;
- Assessment of Technologies and interventions in Health and Rapid Access to Innovation.

In addition to its capacity to maintain a variety of arthropod species in contained environments, VIASEF provides safe housing for in vivo tests of vector-borne pathogens. This allows the development of experimental protocols with multiple research purposes, including:

- The study of vector competence determinants, vector pathogen interactions and immunological factors.
- Testing of new pharmacological products interfering with the development of the pathogen in the vector.
- Testing new insecticidal and repellent candidates, including novel formulations (e.g. nano-encapsulation); and studies of insecticide resistance.
- Performing bioecological and behavioral studies of arthropod vectors towards the development of innovative non-insecticidal vector control tools.