

Boston University Institutional Biosafety Committee (IBC) March 19, 2024 Meeting Minutes Location: Zoom and/or by phone Start time: 12:00 PM End time: 1:35 PM

Members Present:	R. Ingalls, B. Slack, E. Muhlberger, R. Davey, I. Afasizheva, V. Gouon-Evans, P. Liu (joined
	12:08 PM), T. Winters, R. Morales, N. Dey, S. Niemi, J. Keeney, V. Britton (joined 12:07
	PM), S. Ghosh
Guests Present:	J. Connor, A. Ellis, A. Ahmad, A. Broos-Caldwell, P. Richmond, J. Wood, M. Fitzgerald, T.
	Killeen
Staff Present	C. McGoff, L. Campbell

 Review of February 16, 2024 IBC Meeting Minutes (R. Ingalls) No concerns were voiced. Motion: Approved For: 12; Against: 0; Abstain 0; Absent: 2

II. Chair's Report: Nothing to report

III. New Business:

- A. IBC Office Updates: Members were presented with an overview of the changes made to the RIMS online application form.
- B. Incident Report: Nothing to report.
- C. Review of Research Occupational Health Program (ROHP) Report: Nothing to report.
- D. Environmental Health and Safety (EHS) Report: Nothing to report.

IV. Protocol Review

1. rDNA/Bhz – Annual Renewal

BUA	(PI)	Title	BSL	ABSL	Campus
2442	Mohsan Saeed	Investigating the role of viral proteins in pathogenesis	disease 3	N/A	BUMC
Primary	Reviewer: John Conn	or Sec	ondary Reviewer:	Ron Morale	S
••	le NIH Guidelines: Se ents described in this	ee sections III-D-1-b, III-D-2-a, III-D-3-b; App protocol.	endix B-III-D and	G-II-C for	
•		mittee decided to defer the decision on th	· ·	•	

containment without hearing clarification from the NEIDL EHS and NEIDL management about why this move is necessary and how this move is going to impact the work of other PIs who also work in the same BSL-3 labs. The committee recommended that all of the changes pertaining to moving poliovirus work to BSL3 lab be removed so that the regular annual renewal of the protocol can be approved without any concerns. The committee will inform the PI once the aforementioned clarification is available, at which time the PI will be asked to submit an IBC amendment which should include plan to move poliovirus work in BSL3 lab. In the meantime, the following will be communicated to the PI:

- In the context of inactivation procedure to be used, it is stated that "All of the procedures will be performed according to the following recently submitted SOPs.....". These SOPs are now approved. Please provide specific SOP numbers.
- Typo 'Fir' should be 'For'.
 It is stated that 'I am also encouraging my lab members to receive a bivalent SARS-CoV-2 vaccine.' Please update this to state whether there are new procedures or if the vaccination remains ongoing? Clarify if this is relevant to the current state of circulating virus and the strains used in the laboratory.
- Section VIII.6 Describe safety precautions to be practiced for using sharps.

• Please comment on the following concern: Mohoney polio strain listed in the Hazardous biological agent section is PV type 1, which has not been eradicated. Are labs allowed to use the eradicated strains anymore?

Motion: Conditional Approval (Administrative Review) For: 14 Recuse: 0 Against: 0 Abstain: 0 Absent: 0

2. rDNA – New Application

BUA	(PI)	Title		BSL	ABSL	Campus	
2636	Miguel Jimenez	Project 1: Microbial devices for sensing and		1	N/A	CRC	
		actuation					
Primary Reviewer: Rob Davey Secondary Reviewer: Jim Keeney							
Applicab	Applicable NIH Guidelines: Section III-F-8						
Meeting	Meeting Comments: The goal is to make sensors based on microorganism-electronic interfaces. Bacteria and yeast						

are modified to express sensor proteins as well as markers such as luciferase or fluorescent proteins and incorporated into electronic systems such that their reaction to environmental changes can be detected. The main risk components are recombinant DNA-based and production of recombinant organisms. The bacteria and yeast will be *E. coli* and *S. cerevisiae* respectively. To mitigate the risk they will decontaminate liquids with bleach (10% final) for at least 30 minutes, and will use biohazard boxes for solids. PPE used are lab coats, disposable gloves and safety glasses, which are appropriate for the work. Sharps are disposed on in sharps containers. Recombinant plasmids will be made for expression of fluorescent proteins (GFP) as well as transcription factors or signal transduction proteins using standard E. coli and yeast vectors. These include tetR – tetracycline resistance and estrogen receptor and GPCRs from humans. No mammalian cell culture or animal work was proposed. Personnel are trained by the PI, who has appropriate experience.

BUA Site Assessment: This is a new lab which is in the process of getting set up. No biosafety cabinet will be used in
this protocol.. The lab freezers are in lab # which is contiguous to lab # Sharps containers available.Motion: ApproveFor: 14Recuse: 0Against: 0Abstain: 0Absent: 0

3. rDNA/Bhz – Three-Year Renewal

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BUA	(PI)	Title		BSL	ABSL	Campus		
1605	David Levin	Cell Wall Integrity Signaling in Yeast		1	N/A	BUMC		
Primary Reviewer: Valerie Gouon-Evans Secondary Reviewer: Bob Timmerman								
Applicable NIH Guidelines: Section III-F-3, III-F-8, App E-II-A, E-III-A								

Meeting Comments: The goal of the project is to understand how yeast cells maintain the structural integrity of their cell walls to identify molecular targets for the development of antifungal agents that display selective toxicity against fungal cells. Model genetic organisms, such as baker's yeast, *Saccharomyces cerevisiae*, will be used to study the molecular mechanisms of stress signaling, especially the mechanisms by which various stress signals are transmitted to Stress Activated Protein Kinases (SAPK) and downstream pathways. Some of the stressors used include arsenate, arsenite, acetic acid, genomic damage, and curcumin. Genetic modifications in yeast cells will be done by variety of methods including gene knock-outs, gene replacements, plasmid-based gene expression, plasmid library screens for complementation or suppression, mutant screens and selections, etc. They use nonpathogenic E. coli to clone and express recombinant proteins and to make variety of plasmids for introducing foreign genes into yeasts. The lab generates point mutants and deletion mutants of various genes using PCR-mediated random and directed mutagenesis. The list of genes mutated in the past are provided and indicate that other genes to be mutated will depend on the results of the screens. The following will be communicated to the PI:

- Check vigorous mixing, blending
- Check Laboratory Coats
- Remove N95. It is not required for the proposed procedures.
- Please add a sentence for the liquid biohazard disposal that all liquid wastes will be treated with bleach at a final concentration of 10% for a minimum of 30 minutes before disposal down the drain.

BUA Site Assessment: Add procedure: Vigorous mixing/blending Add PPE: Laboratory coats, Remove PPE: N-95respirator. BSC not needed for proposed work. Roomcan be removed from the protocol as it is an autoclave and
washing room. No biohazardous material transportation is anticipated in this protocol.Motion: Conditional Approval (Administrative Review)For: 14Recuse: 0Against: 0Abstain: 0Absent: 0

4. rDNA/Bhz – Amendment

BUA	(PI)	Title		BSL	ABSL	Campus		
2566	Andreea Bujor	Biorepository in systemic sclerosis		2	N/A	BUMC		
		Systemic sclerosis research studies						
Primary	Reviewer: Inna Afasi	izheva	Secondary Revi	econdary Reviewers: Valerie Gouon-Evans				
Applicat	Applicable NIH Guidelines: Sections III-D-1-a, III-E-1							
Meeting	Comments: The goa	al of this study is to investigate how im	mune cells move	e through	the skin l	ayer in systemic		
sclerosis	(Scleroderma) and t	o cellular signaling pathways that con	tribute to the ab	normal in	nmune ce	ll function.		
Specifica	Specifically they use biological samples from scleroderma patients including heart and lung tissues. This amendment							
to add n	ew procedures by w	hich the group will explore the role of	specific transcrip	otion fact	or in Scler	oderma. They		
will lent	will lentivirus vectors to manipulate expression of this transcription factor in human cancer cell lines and try to							
correlat	e their findings with	what is observed in clinical samples. T	he following will	be comm	nunicated	to the PI:		
	Since the protocol p	a protocol now involves rDNA work (lentivirus work) a "Not applicable to this project" response for						

- Since the protocol now involves rDNA work (lentivirus work), a "Not applicable to this project" response for rDNA training is not appropriate for any of the listed personnel (such as a second or second o
- All listed members must complete the rDNA/IBC policy training in BioRAFT, even when someone will not be doing any rDNA work. This is mandatory.
- VIII. 1: add "pipetting infectious liquid
- VIII.5: Please provide a recent certification date of the biosafety cabinet.
- VIII.8: Please list the disinfectants to be used.
- VIII.10 and VIII.11: please answer (How and where are biohazardous materials stored and transported such as human cancer cell line, lentivirus stock)
- Please add lentivirus in the hazardous biological agent list
- Please update IRB approval dates: : 4/24/2025; : 3/14/2025.
- Replace the current statement in the Applicable NIH guidelines question with "Section III-D-1-a, and III-E-1."

Motion: Conditional Approval (Administrative Review) For: 14 Recuse: 0 Against: 0 Abstain: 0 Absent: 0

5. rDNA/Bhz – Amendment

BUA	(PI)	Title		BSL	ABSL	Campus	
2350	Robert Davey	Viruses pseudotyped with hemorhagic fever virus			N/A	BUMC	
		glycoproteins and virus-like particles for study of					
		infection mechanism.					
Primary Reviewer: Elke Muhlberger Secondary Re				viewer: Saj	al Ghosh		
Applicab	Applicable NIH Guidelines: III-D-1-a, III-D-2-a, III-D-3-a, III-E-1						
Meeting	Meeting Comments: The original goal of this protocol is to make pseudotyped viruses or virus like particles for their						

use as reporter systems to detect inhibitors of virus entry. The viruses are made by expression of virus proteins in cells. Each virus generated is replication defective, only able to infect cells but unable to assemble new viruses. A marker of infection is encoded within the recombinant virus genome such as a fluorescent protein or luciferase enzyme. In this amendment they propose to work with plasmids containing segments of filovirus, arenavirus and bunyavirus genomes. The virus types will be limited to arenaviruses, which include New World (Junin and Machupo) and Old World (Lassa) viruses as well as Crimean Congo Hemorrhagic fever virus (CCHFV) that have multi-segmented genomes, as well as parts of Ebolaviruses where the genome is normally on one segment. In each case, plasmids containing each segment will be cultivated in *E. coli* as plasmids. They will then be mutated to place in point mutations that are expected to cause reductions in virus replication fitness to understand replication mechanism. For this protocol, at all times, the genome plasmids are handled independently of each other, never in combination and support plasmids (those needed to make the virus RNA replication complex) are kept away from each genome fragment and away from mammalian tissue culture. NIH has approved PI's request to perform these additional arenavirus genome cloning work in BSL2 lab. The committee requested few minor clarification on the research plan. The following will communicated to the PI:

- This amendment covers plasmid preparations of viral genome fragments and introducing mutations in these plasmids. It is not clear why the PI added this sentence: Each change would not be expected to increase virus virulence and if such were identified, we would stop work and notify the IBC and EHS, as detailed in our BSL4 protocols." This refers to viruses generated with these constructs, but these are not part of this protocol. Needs clarification.
- It is stated that "We propose to perform the work at BSL2 in a space that is distinct from our tissue culture room with a normally closed door and engineering controls separating the spaces." This room is different from the full-length clone room, correct? Please clarify.
- It might be helpful to remove the generation of full-length filovirus plasmids from this protocol and add that this work is covered by IBC protocol (PI (PI (PI))) the reason being that there are NIH regulations in place for handling full-length filovirus and henipavirus clones. Members of the PI's lab are listed on protocol
- The PI may wish to add that *E. coli* K12 bacteria will be used.
- Section VIII, 1: Check "Culture stirrers, shakers" and "Plating, colony counting".
- Section VIII, 6: If sharps are used to cut out agarose gel fragments, this should be added to section.
- rDNA section The PI states that fragments from "other filoviruses" will be generated. IBC recommends naming the other filoviruses.
- rDNA section, eukaryotic host strains: It is stated that some cell lines will be used "as recipients of recombinant viruses". Since no recombinant full-length virus is being put in cell culture in this protocol, clarify what this sentence mean. This sentence should be removed if it relates to BSL4 work.
- rDNA section, 8 and 11: Please confirm that no DNA fragments >200 nts will be ordered to generate viral fragments.

PI recused himself from approval discussion and voting.

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Motion: Conditional Approval (Administrative Review)	For: 13	Recuse: 1	Against: 0	Abstain: 0	Absent: 0

6. rDNA/Bhz – Amendment

BUA	(PI)	Title	E	BSL	ABSL	Campus							
2562	Philip Strandwitz	Discovery and development of thera	pies and 2	2	N/A	BUMC							
		consumer products derived from the	e human										
		microbiome											
Primary I	Reviewer: Robin Inga	alls	Secondary Review	wer: Inn	a Afasizhev	/a							
Applicab	le NIH Guidelines: S	ections III-D-1-a and III-D-2-a											
Meeting	Comments: The goa	l of this protocol is to identify as man	y as possible indiv	idual hu	man gut ba	acterium							
species t	hat are believed to o	contribute to good health. They will sc	reen diverse pane	els of hur	nan gut ba	icteria for							
activity a	gainst major disease	e targets. Once they have an understa	nding of which ba	cteria in	fluence a s	pecific disease							
target, th	ney plan to engineer	that bacteria to be able to use it as p	obiotic and for co	mmercia	al develop	ment. In this							
amendment they propose for the first time, to add a recombinant DNA procedure by which they will be able to													
delete sp	delete specific unwanted gene from the bacteroides species. The process includes plasmid manipulation by												
recombir	recombinant DNA technology, introduction of the plasmid into <i>E. coli</i> and subsequent transfer of the plasmid to the												
bacteroides species by bacterial conjugation. Specific cloning technology to be used in the protocol, will allow the													
bacteroid	bacteroides to expel the desired gene from its genome. The experimental procedures are described in detail and they												
do not a	opear to involve risk	of exposure to hazardous materials. T	he following will l	be comn	nunicated	do not appear to involve risk of exposure to hazardous materials. The following will be communicated to the PI:							

- PI must complete rDNA/IBC Policy training in BioRAFT. All members of a protocol involving rDNA work must also complete the rDNA/IBC policy training.
- Please correct the typo on the sentence in the laboratory procedure section "However, since pARC [luck] a Bacteroides origin of replication.." This should be 'lacks'.
- Provide recent Biosafety Cabinet certification date.

Motion: Conditional Approval (Administrative Review)	For: 14	Recuse: 0	Against: 0	Abstain: 0	Absent: 0
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V. List of Protocols reviewed by DMR (not discussed in the meeting)

A list of protocols that were reviewed by DMR was displayed in the meeting.