

Biological Toxin Safe Work Practices

I. INTRODUCTION

Biological toxins are poisonous substances produced by certain microorganisms, animals, and plants. Examples of toxins of biological origin include diphtheria toxin, tetrodotoxin, pertussis toxin, botulinum neurotoxin, snake venom toxins, conotoxin and Ricin. Although toxins are derived from biological materials, they do not replicate and are therefore not considered infectious. However, they may be extremely toxic in very small quantities and must be managed like hazardous chemicals in the workplace. Controls must be in place to ensure staff is protected from exposure. The routes of exposure include inhalation, eye, nose and mucous membrane contact, percutaneous, and skin absorption depending on the diluents used. The main issues of concern in the laboratory are accidental exposures to toxin caused by contact with the mouth, eye, skin and mucous membranes, inhalation of toxin powder or aerosol inadvertently generated, or by needlestick incidents.

Work with toxins of biological origin must be included in your laboratory-specific Chemical Hygiene Plan. Documented toxin-specific hazard training and training on the laboratory-specific standard operating procedures (SOP) is required for all laboratory personnel prior to starting work. The training must include, but is not limited to, the health and physical hazards of the toxin, signs and symptoms associated with exposure, appropriate work practices, personal protective equipment, and emergency procedures.

Some toxins of biological origin are considered Select Toxins, which the US Departments of Health and Human Services (HHS) and Agriculture (USDA) have determined to have the potential to pose a severe threat to public health and safety, to animal or plant health, or to animal or plant products. The Centers for Disease Control and Prevention (CDC) in their Select Agent Program regulate the possession, use, and transfer of these specific biological agents and toxins. Research work with CDC listed Select Toxins may have additional safety and security requirements including registration with the UW Select Agent Program and the CDC.

II. LABORATORY PLANNING AND PREPARATION FOR USE

- Certain toxin forms and toxins used in minimal quantities may be excluded from the requirements of Select Agent Regulations. The CDC maintains a current list of Select Toxins, exclusions, and exempt quantities on the <u>Federal Select Agent Program website</u>. For work with <u>select toxins</u> at or above the <u>permissible amounts</u>, you are required to enroll in the UW Select Agent Program and the Federal Select Agent Program. Email <u>uwsa@uw.edu</u> to find out if you need to enroll.
- Research involving botulinum neurotoxin *in any quantity* requires review for Dual Use Research of Concern (DURC). More information is available on the <u>EH&S DURC</u> <u>webpage</u>.

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- 3. Develop a written laboratory-specific SOP specific to the toxin being used. A template <u>Biological Toxin SOP</u> with training documentation form is available. A specific <u>diphtheria</u> <u>toxin SOP</u> template is also available.
- 4. Provide and document hazardous chemical training and specific toxin SOP training to personnel working with toxins and any other personnel authorized or required to be in the laboratory during toxin work. A sample training documentation form is included in the template SOP referenced above.
- 5. Ensure the toxin Material Safety Data Sheet/Safety Data Sheet (MSDS/SDS) is available to staff at all times and that the toxin inventory for the laboratory is entered into the <u>UW MyChem system</u>.
- 6. Designate toxin storage area in a locked container (freezer, refrigerator, cabinet or other container) in a secure location.
- 7. Designate a laboratory, work space, and certified biological safety cabinet (BSC), fume hood, glove box or other approved containment for toxin work. The laboratory facilities required may vary based on the level of hazard posed by the specific toxin and the procedures being performed. Work with Select Toxins may require rooms with controlled access.
- 8. Prepare a door sign stating "Toxins in Use Authorized Personnel Only."
- 9. Post the EH&S Exposure Response Poster and EH&S Spill Response Poster in the laboratory.
- 10. If possible, do not work with toxin in solid or powder form. If it is necessary to purchase it in powder or solid form, purchase pre-diluted or pre-weighed toxin in the minimum quantity needed to perform work. Additional precautions may be needed if working with powder or solid toxin.
- 11. Determine the appropriate chemical and/or physical inactivation method(s) for the specific toxin (refer to toxin inactivation Section VIII). Ensure supplies for inactivation of toxin are available.
- 12. Ensure supplies for spill cleanup are appropriate for the specific toxin, maintained in a clearly marked spill cleanup kit and readily available in the laboratory.

III. ENGINEERING CONTROLS

- 1. Designate a certified BSC, fume hood, glove box or other approved containment. Do not use a laminar flow hood or cabinet for toxin work. Consider the properties of the specific toxin and procedures when selecting a containment device.
- 2. In-line HEPA filters are required if vacuum lines are used with toxin.
- 3. If centrifuging materials containing toxin, centrifuge safety cups or sealed rotors must be used and the outside surfaces routinely decontaminated. Open the sealed cups or rotors inside containment.

IV. PERSONAL PROTECTIVE EQUIPMENT (PPE)

- 1. Wear safety glasses with side shields or goggles.
- 2. Wear a laboratory coat with long sleeves, smock, apron, or coveralls. Consider using disposable PPE.

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- 3. Wear gloves that are impervious to the toxin as well as the diluent. Double gloving is recommended. Change gloves immediately if contaminated, torn, or punctured and dispose immediately after removal.
- 4. Wear face protection, such as a face shield, when splash/splatter is possible.
- 5. Respiratory protection (requires enrollment in UW's respirator program) may be required if an airborne hazard is present when work is done outside of approved containment. Refer to the <u>EH&S Respiratory Protection webpage</u>.

V. TOXIN USE PRACTICES (reconstitution, dilution, administration)

- 1. Post sign on room door when toxins are in use stating "Toxins in Use Authorized Personnel Only."
- 2. Work with toxins in designated rooms at pre-determined bench areas.
- 3. Biosafety level 2 (BSL-2) practices are appropriate for most toxin work. However, some toxins or procedures may require additional BSL-3 practices.
- 4. Work with toxin in a BSC, fume hood, glove box or other approved containment.
- 5. Transport toxins only in labeled, leak/spill-proof, non-breakable secondary containers.
- 6. Perform preparations over plastic backed absorbent pads. Dispose of pads after completion of tasks or immediately upon contamination.
- 7. Utilize safe sharps procedures (i.e., sharps container in the immediate vicinity). Needle locking syringes or disposable syringe needle units are recommended and should be disposed of promptly after use.
- 8. Restrain or anesthetize animals when possible.
- 9. Decontaminate containers before they are removed from the fume hood, BSC, or glove box. Also decontaminate the exterior of the closed primary container and place it in a clean secondary container.
- 10. Decontaminate the BSC or approved containment and all surfaces used upon completion of tasks with appropriate inactivating agent and contact time.
- 11. All potentially contaminated disposable items (such as gloves used in preparation) must be placed in a hazardous waste bag and autoclaved before disposal.
- 12. Wash hands upon completion of tasks.

VI. TOXIN SPILL CLEANUP

Toxin spills must be cleaned up immediately by properly protected and trained personnel. Follow the instructions on the <u>EH&S Spill Response Poster</u> as shown below. More detailed instructions follow.

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SPILL RESPONSE



CALL 911 FOR ANY LIFE THREATENING EMERGENCY IF EXPOSED, FOLLOW THE EXPOSURE RESPONSE POSTER

S.W.I.M. FOR ALL SPILLS

- **S: Stop** the spill. Cover with absorbent material.
- **W:** Warn others. Alert people in the immediate area of the spill.
 - **I:** Isolate the spill and secure the area. Close doors if possible.
- M: Minimize your exposure by wearing PPE and avoiding contact, inhalation or ingestion. Vacate the area if necessary. Wash hands after handling spill materials.

RADIOLOGICAL SPILLS

- Utilize time, distance and shielding to prevent exposure.
- Cover with absorbent material.
- Wear gloves and use tongs/scoop to collect contaminated material as radioactive waste.
- Call UW Radiation Safety at 206.543.0463. If office closed, call 911.
- Notify your supervisor.

CHEMICAL SPILLS

- *If exposed, use the eye wash or safety shower for 15 minutes.*
- Large spills: Pull the fire alarm and evacuate. EH&S can arrange for hazardous spill cleanup at the lab's expense.
- **Small spills:** Trained personnel familiar with the chemical should use the lab's spill kit.
- Staff must protect themselves from skin, eye and respiratory hazards by using personal protective equipment (PPE) during cleanup.

EH&S chemical spill assistance is available 24/7:

- During business hours (Monday-Friday 8 a.m. to 5 p.m.), call **206.543.0467**.
- After business hours, call 206.685.UWPD (8973) to reach EH&S staff on call.

BIOHAZARDOUS SPILLS

- Cover the spill with paper towels or absorbent material.
- Pour freshly prepared 10% bleach around the spill and allow to flow into spill.
- After 30 minutes of contact time, wipe up and dispose of as biohazard waste.
- Repeat procedure.

If spill contains recombinant nucleic acids, notify EH&S Biosafety as soon as possible at **206.221.7770**.

Report all spills within 24 hours via UW OARS: <u>https://oars.ehs.washington.edu</u>

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www.ehs.washington.edu

View an accessible version of the <u>Spill Response Poster</u> on the EH&S website.



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Liquid Spills

- 1. The required PPE for cleaning up spills includes a lab coat or smock, goggles, and two pairs of nitrile gloves.
- 2. For chemical inactivation, cover spill with absorbent paper towels and inactivate by applying the appropriate chemical inactivating agent starting at the perimeter and working toward the center, allowing prescribed contact time before clean up. Clean the spill area with inactivating agent, allowing prescribed contact time, then soap and water. The inactivated spill waste can be double bagged and disposed of in regular trash. Toxin inactivation information is in section VIII below.
- 3. For physical inactivation use absorbent paper towels to wipe up liquid. Place waste in hazardous waste plastic bag and autoclave. Clean the spill area with inactivating agent, allowing prescribed contact time, then soap and water. The inactivated spill waste can be double bagged and disposed of in regular trash. Toxin inactivation information is in section VIII below.

Powder Spills inside a BSC or containment

- 1. The required PPE for cleaning up spills includes a lab coat or smock, goggles, and two pairs of nitrile gloves.
- 2. Gently cover powder spill with dampened absorbent paper towels to avoid raising dust.
- 3. For chemical inactivation, apply the appropriate chemical inactivating agent starting at the perimeter and working toward the center, allowing prescribed contact time before clean up. Clean the spill area with inactivating agent, allowing prescribed contact time, then soap and water. The inactivated spill waste can be double bagged and disposed of in regular trash. Toxin inactivation information is in section VIII below.
- 4. For physical inactivation, use dampened paper towels to wipe up spill. Place waste in hazardous waste plastic bag and autoclave. Clean the spill area with inactivating agent, allowing prescribed contact time, then soap and water. The inactivated spill waste can be double bagged and disposed of in regular trash. Toxin inactivation information is in section VIII below.

Powder spills outside of a BSC, fume hood, glove box or approved containment

- 1. Remove all personnel from the laboratory and restrict access; do not attempt to clean up the spill.
- As soon as possible report the spill by notifying EH&S (EH&S business hours 206-543-0467, outside business hours 911); tell them that a spill has occurred, and you need EH&S to obtain a spill cleanup contractor.
- 3. Be prepared to provide the following information:
 - Name and phone number of knowledgeable person that can be contacted
 - Name of toxin, concentration and amount spilled, liquid or solid type spill

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- Number of injured, if any (refer to Section VII Acute Exposure)
- Location of spill

This information can also be used in reporting to the Emergency Department (ED) after potential exposure. The involved person or supervisor is required to complete and submit the <u>Online Accident Reporting System (OARS)</u> form to EH&S within 24 hours of any spill incident. For questions on spill cleanup, contact EH&S spill consultants at 206-543-0467 for guidance.

VII. ACUTE EXPOSURE

Follow the steps on the EH&S Exposure Response Poster below for any exposures to biotoxins.

Note: For diphtheria toxin exposures, refer to the <u>EH&S SOP template for Diphtheria Toxin</u> for specifically required Emergency Department procedures.

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EXPOSURE RESPONSE

for biological, chemical, or radiological exposures

CALL 911 FOR ANY LIFE THREATENING EMERGENCY

1. PERFORM FIRST AID Needlestick, puncture or sharps Wash thoroughly for 15 minutes with warm water and injury, or animal bite/scratch sudsing soap. **Eye exposure** Use emergency station to flush eyes for 15 minutes while holding eyes open. Skin exposure • Radioactive: Survey skin and wash until the count rate cannot be reduced further. Stop if skin becomes irritated. • Chemical: Wash with tepid water for 15 minutes. • Hydrofluoric acid: Wash for 5 minutes, then apply calcium gluconate gel to skin. • Biological: Wash with sudsing soap and water for 15 minutes. Inhalation or ingestion • Move out of the contaminated area and seek fresh air. • Do not induce vomiting unless instructed to do so. • Radioactive: Blow nose into clean tissue and survey for contamination. 2. GET MEDICAL HELP For radiological exposure • Call Radiation Safety at 206-543-0463. or emergency: • Call 911 if office closed. · Provide the radionuclide, estimated amount and time since exposure. For chemical exposure • Call 911 and follow the instructions given. or emergency: • Provide the chemical name, concentration, time since exposure and Safety Data Sheet (SDS). For biological and • Call the Employee Health Center at 206-685-1026. all other exposures: • Harborview sites call 206-744-3081. • If closed, call **911** and follow the instructions given. For all exposures: • Notify your supervisor. Secure the area before leaving. **3. REPORT THE INCIDENT** For hospitalization, fatality, Notify EH&S immediately after performing first aid and getting or recombinant nucleic acid medical help: exposure: • Call the EH&S main phone line at 206-543-7262. If closed, call 206-685-UWPD(8973) to reach EH&S staff on call. All incidents and near misses: Submit a report via the UW Online Accident Report (OARS) within 24 hours at https://oars.ehs.washington.edu. September 2019

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View an accessible version of the <u>Spill Response</u> Poster on the EH&S website.

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VIII. INACTIVATION AND DISPOSAL

According to the CDC, inactivation of a biotoxin means to render the toxin non-functional so that it is no longer capable of exerting its toxic effect. This is different from inactivation of biological agents, which renders the agent non-viable, or no longer capable of growing, replicating, infecting, or causing disease. Inactivation methods used for biotoxins must be specific for the toxin, published and validated, or developed and validated with thorough testing. Note that disinfecting solutions and products may not inactivate biotoxins.

- 1. Inactivate any waste toxin chemically or physically (usually autoclaving) before disposal or given to EH&S for disposal.
- 2. Place any used PPE and spill cleanup debris in a hazardous waste plastic bag and autoclave.
- 3. For mixed waste (i.e., toxin waste mixed with radioactive waste) consult EH&S Radiation Safety at 206-543-0463 for disposal instructions.
- 4. If in-lab inactivation is not possible for some toxin waste, manage waste as hazardous chemical waste. Be aware that some form of treatment in the lab may be required before collection as chemical waste. Contact EH&S Environmental Programs Office at 206-616-5835 for disposal instructions. Refer to the EH&S Chemical Waste Disposal webpage to request collection of hazardous chemical waste.
- 5. Dispose of liquid inactivated biotoxin waste within a pH range of 5.5 to 12 down the regular sewer drain.
- Refer to the information below in Tables 1 and 2 on inactivation of selected toxins, which is taken directly from the publication Biosafety in Microbiological and Biomedical Laboratories (BMBL) 5th Edition. Centers for Disease Control and Prevention. Appendix I: Guidelines for Work with Toxins of Biological Origin.
- If using bleach solutions, prepare fresh daily for inactivation of biotoxins and decontamination of surfaces. Undiluted, commercially available bleach solutions typically contain 3 – 6% (w/v) NaOCI (sodium hypochlorite).
- 8. Since diphtheria toxin is not included in the BMBL tables, a review was made of inactivation methods for Diphtheria Toxin at various research institutions. The most common physical inactivation method was steam autoclaving at 121°C for 60 minutes. Although no consensus was apparent for a specific chemical inactivation agent and concentration, the commonly used chemicals included 1% NaOCI, 10% bleach, 1N NaOH, and combinations of NaOCI and NaOH. A 30-minute contact time was allowed to complete inactivation.

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ΤΟΧΙΝ	STEAM AUTOCLAVE	DRY HEAT (10 MIN)	FREEZETHAW	GAMMA IRRADIATION
Botulinum neurotoxin	Yesa	> 100ºCb	Νοα	Incompleted
Staphylococcal Enterotoxin	Yese	> 100ºC; refoldsf	Nog	Incompleteh
Ricin	Yesi	> 100ºCi	Noj	Incompletek
Microcystin	Νοι	> 260ºCm	Non	ND
Saxitoxin	Νοι	> 260ºC m	Non	ND
Palytoxin	Νοι	> 260ºCm	Non	ND
Tetrodotoxin	Νοι	> 260ºCm	Non	ND
T-2 mycotoxin	Νοι	> 815ºCm	Non	ND
Brevetoxin (PbTx-2)	Νοι	> 815ºCm	Non	ND

IX. TABLE 1: Physical Inactivation of Selected Toxins

Table 1 Notes: ND indicates "not determined" from available decontamination literature.

^a Steam autoclaving should be at \geq 121°C for 1 h. For volumes larger than 1 liter, especially those containing *Clostridium* botulinum spores, autoclave at \geq 121°C for 2 h to ensure that sufficient heat has penetrated to kill all spores.

^b Exposure to 100°C for 10 min. inactivates BoNT. Heat denaturation of BoNT as a function of time is biphasic with most of the activity destroyed relatively rapidly, but with some residual toxin (e.g., 1-5%) inactivated much more slowly.

^c Measured using BoNT serotype A at -20°C in food matrices at pH 4.1-6.2 over a period of 180 days.

^d Measured using BoNT serotypes A and B with gamma irradiation from a Co source.

^e Protracted steam autoclaving, similar to that described for BoNT, followed by incineration is recommended for disposal of SEcontaminated materials.

Inactivation may not be complete depending upon the extent of toxin re-folding after denaturation. Biological activity of SE can be retained despite heat and pressure treatment routinely used in canned food product processing.

^g SE toxins are resistant to degradation from freezing, chilling or storage at ambient temperature. Active SEB in the freeze-dried state can be stored for years.

h. References 15,16 in BMBL

Dry heat of \geq 100°C for 60 min in an ashing oven or steam autoclave treatment at >121°C for 1 h reduced the activity of pure ricin by >99%. Heat inactivation of impure toxin preparations (e.g. crude ricin plant extracts) may vary. Heat-denatured ricin can undergo limited refolding (<1%) to yield active toxin. Ricin holotoxin is not inactivated significantly by freezing, chilling or storage at ambient temperature. In the liquid state with a preservative (sodium azide), ricin can be stored at 4°C for years with little loss in potency.

Irradiation causes a dose-dependent loss of activity for aqueous solutions of ricin, but complete inactivation is difficult to achieve; 75 MRad reduced activity 90%, but complete inactivation was not achieved even at 100 MRad. Gamma irradiation from a laboratory Co source can be used to partially inactivate aqueous solutions of ricin, but dried ricin powders are significantly resistant to inactivation by this method.

Autoclaving with 17 lb pressure (121-132 °C) for 30 min failed to inactivate low molecular weight (LMW) toxins. All burnable waste from LMW toxins should be incinerated at temperatures in excess of 815°C (1,500 °F).

"Toxin solutions were dried at 150°C in a crucible, placed in an ashing oven at various temperatures for either 10 or 30 min, reconstituted and tested for concentration and/or activity; tabulated values are temperatures exceeding those required to

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achieve 99% toxin inactivation.

["]LMW toxins are generally very resistant to temperature fluctuations and can be stored in the freeze-dried state for years and retain toxicity.

ΤΟΧΙΝ	NAOCL (30 MIN)	NAOH (30 MIN)	NAOCL + NAOH (30 MIN)	OZONE TREATMENT
Botulinum neurotoxin	> 0.1%a	>0.25 N	ND	Yes
Staphylococcal enterotoxin	> 0.5%c	>0.25 N	ND	ND
Ricin	> 1.0%d	ND	>0.1% + 0.25N e	ND
Saxitoxin	≥ 0.1%e	ND	0.25% + 0.25N e	ND
Palytoxin	≥ 0.1%e	ND	0.25% + 0.25N e	ND
Microcystin	≥ 0.5%e	ND	0.25% + 0.25N e	ND
Tetrodotoxin	≥ 0.5%e	ND	0.25% + 0.25N e	ND
T-2 mycotoxin	≥ 2.5%e,f	ND	0.25% + 0.25N e	ND
Brevetoxin (PbTx-2)	≥ 2.5%e,f	ND	0.25% + 0.25N e	ND

X. TABLE 2: CHEMICAL INACTIVATION OF SELECTED TOXINS

Table 2 Notes: ND indicates "not determined" from available decontamination literature.

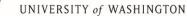
^a Solutions of NaOCI (≥0.1%) or NaOH (>0.25 N) for 30 min inactivate BoNT and are recommended for decontaminating work surfaces and spills of *C. botulinum* or BoNT. Chlorine at a concentration of 0.3-0.5 mg/L as a solution of hypochlorite rapidly inactivates BoNT (serotypes B or E tested) in water. Chlorine dioxide inactivates BoNT, but chloramine is less effective. ^b Ozone (>2 mg/L) or powdered activated charcoal treatment also completely inactivate BoNT (serotypes A, B tested) in water under defined condition.

° SEB is inactivated with 0.5% hypochlorite for 10-15 mi.

^d Ricin is inactivated by a 30 min exposure to concentrations of NaOCI ranging from 0.12.5%, or by a mixture of 0.25% NaOCI plus 0.25 N NaOH.¹⁷ In general, solutions of 1.0% NaOCI are effective for decontamination of ricin from laboratory surfaces, equipment, animal cages, or small spills.

^e The minimal effective concentration of NaOCI was dependent on toxin and contact time; all LMW toxins tested were inactivated at least 99% by treatment with 2.5% NaOCI, or with a combination of 0.25% NaOCI and 0.25N NaOH.
^f For T-2 mycotoxin and brevetoxin, liquid samples, accidental spills, and non-burnable waste should be soaked in 2.5% NaOCI with 0.25% N NaOH for 4 h. Cages and bedding from animals exposed to T-2 mycotoxin or brevetoxin should be treated with 0.25% NaOCI and 0.025 N NaOH for 4 h. Exposure for 30 min to 1.0% NaOCI is an effective procedure for the laboratory (working solutions, equipment, animal cages, working area and spills) for the inactivation of saxitoxin or tetrodotoxin.
Decontamination of equipment and waste contaminated with select brevetoxins has been reviewed.

Alternate methods of chemical decontamination: 1 N sulfuric or hydrochloric acid did not inactivate T-2 mycotoxin and only partially inactivated microcystin-LR, saxitoxin, and brevetoxin (PbTx-2). Tetrodotoxin and palytoxin were inactivated by hydrochloric acid, but only at relatively high molar concentrations. T2 was not inactivated by exposure to 18% formaldehyde plus methanol (16 h), 90% freon-113 + 10% acetic acid, calcium hypochlorite, sodium bisulfate, or mild oxidizing.¹⁷ Hydrogen peroxide was ineffective in inactivating T-2 mycotoxin. This agent did cause some inactivation of saxitoxin and tetrodotoxin, but required a 16 h contact time in the presence of ultraviolet light.



XI. TRANSFER OF SELECT TOXINS

Documentation is required for any transfer of the following Select Toxins shown below in any amount (intramurally or extramurally), to any entity or individual. If you plan to transfer any of these Select Toxins you must complete a <u>UW Select Toxin Transfer Due Diligence Form</u> and submit a copy to the UW Select Agent Program prior to the transfer at <u>uwsa@uw.edu</u>. It is the responsibility of the Principal Investigator transferring the Select Toxin to perform due diligence and to keep records for three years. Contact the UW Select Agent Program at <u>uwsa@uw.edu</u> for questions.

- 1. Abrin
- 2. Botulinum neurotoxins
- 3. Short, paralytic alpha conotoxins
- 4. Diacetoxyscirpenol (DAS)
- 5. Ricin
- 6. Saxitoxin
- 7. Staphylococcal Enterotoxins (Subtypes A, B, C, D, and E)
- 8. T2 toxin
- 9. Tetrodotoxin

XII. RESOURCES

EH&S CONTACTS:

- For biotoxin safety consultation, contact EH&S Biological Safety at ehsbio@uw.edu or 206-221-7770.
- For spills, contact EH&S Chemical Spill Advice by calling 206-543-0467.
- For disposal, visit <u>EH&S Chemical Waste Disposal</u> or email <u>chmwaste@uw.edu</u> or call 206-616-5835.
- For toxin waste mixed with radioactive waste, contact EH&S Radiation Safety at 206-543-0463 or radsaf@uw.edu.

SOP TEMPLATES:

- <u>Biological Toxin SOP template</u>
- <u>Diphtheria Toxin SOP template</u>

FORMS:

- <u>UW Select Toxin Transfer Due Diligence Form</u>
- Biological Toxin Checklist

REFERENCE:

• **Biosafety in Microbiological and Biomedical Laboratories (BMBL)** 5th Edition (December 2009). Centers for Disease Control and Prevention. Section VIII-G Toxin Agents, Appendix F: Select Agents and Toxins, and Appendix I: Guidelines for Work with Toxins of Biological Origin. Available <u>online</u>.