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NERVE AGENT INFORMATION FOR EMERGENCY MEDICAL SERVICES AND HOSPITALS

Meticulous attention to standard protocols for personal protection, recognizing toxidromes, and treating patients continues to be the best way to prepare for and respond to chemical agent exposures

Purpose

This document provides a quick refresher on standard protocols for recognizing, treating, and protecting yourself from nerve agent exposures. Comprehensive follow-up guidance for Law Enforcement, Fire, EMS, HazMat, and Hospital-Based First Receivers incorporating lessons learned and best practices from the recent United Kingdom incidents will be forthcoming.

BACKGROUND

Nerve agents are extremely toxic chemical warfare agents. Several nerve agents exist and are generally categorized as either "high volatility" or "low volatility" chemicals, a measure of how likely they are to disperse in air. A high volatility nerve agent (easily dispersed in air) means that the exposure is likely to occur from breathing in its vapors resulting in the rapid onset of symptoms. A low volatility nerve agent (not easily dispersed in air) typically gets absorbed through the skin and has a delayed onset of signs and symptoms. An example of a high volatility nerve agent is sarin, whereas VX is a low volatility agent. In the body, a nerve agent exerts its effects by inhibiting an enzyme (acetylcholinesterase), resulting in acute illness – specifically, cholinergic crisis. Organophosphorus or carbamate pesticides produce similar effects to nerve agents.

SIGNS AND SYMPTOMS OF NERVE AGENT POISONING

Caveat: Poisoned patients <u>may not</u> demonstrate all of these symptoms

- Mouth/Skin: Drooling (Salivation), foaming at the mouth, and excessive sweating
- · Nose/Eyes: Runny nose and watery eyes (Lacrimation) with small (often pinpoint) pupils (Miosis)
- Chest: Cough, chest tightness, difficulty in breathing, wheezing, respiratory failure, "wet" fluid filled lungs
- Abdominal: Urination, Diarrhea, abdominal (Gastrointestinal) cramps, belching, nausea, and/or vomiting (Emesis)
- Mental Status: Confusion, drowsiness, slurred speech, ataxia, unconsciousness, coma
- Muscle/Neurological: Fatigue, weakness, twitching, tremors, cramps, absent reflexes, seizures

Underlined findings = "SLUDGE"- Salivation, Lacrimation, Urination, Diarrhea, Gastrointestinal cramps, Emesis
Other mnemonic used = "DUMBBELS" - Diarrhea, Urination, Miosis/Muscle weakness, Bronchospasm/Bronchorrhea,
Bradycardia, Emesis, Lacrimation, Salivation/Sweating

Clinical Effects of Nerve Agents versus Opioids				
Nerve Agent		Opioid		
Nose	Runny nose	Normal		
Airway	Secretions/drooling/foaming at the mouth	Normal		
Breathing / Respiratory status	Increased work of breathing/chest tightness/wheezing/difficulty in breathing/cough/"wet" fluid filled lungs- more prominent with inhaled exposure; dermal exposure may not cause bronchoconstriction or bronchorrhea			
Heart rate	Slowed	Normal		
Mental Status / Neurological	Slow/unconscious/seizures/confusion/slurred speech/ataxia/coma/absent reflexes/ tremors	Slow or unconscious/coma/seizures		
Eyes	Tearing/small pupils-pinpoint	Small pupils-pinpoint		
Skin	Wet/sweaty/cyanosis	Normal/cyanosis		
Gastrointestinal	Belching/cramps/vomiting/diarrhea	Normal		
GU	Urination	Normal		
Muscles	Fatigue/weakness/twitching/ cramps	Normal		

A KEY DISTINCTION BETWEEN NERVE AGENT POISONING AND OPIOID POISONING IS "SLUDGE" OR "DUMBBELS."

DETECTION (IF YOU STRONGLY SUSPECT A NERVE AGENT)

- Contact HazMat or special operations teams
- · Notify the local FBI Field Office WMD coordinator

PERSONAL PROTECTIVE EQUIPMENT (PPE)

- Emergency responders should have the proper training and education to work with hazardous materials.
- Those providing or assisting with patient care including decontamination should follow institutional policy for a chemical incident, wearing a recommended chemical protective suit, gloves, boots, and respiratory protection to prevent any secondary exposure from patients or objects.
- After patient decontamination is complete, providers should wear a gown and a double layer of nitrile gloves during patient contact.

PATIENT DECONTAMINATION

- A person potentially exposed to a nerve agent should be decontaminated whether they develop signs of acute illness or not.
- Removal of clothing is a vital step to reduce ongoing and secondary exposure. Responders should pay particular attention to the risk of secondary exposure during clothing removal. Double bagging removed clothing is ideal.
- Wiping skin with a paper towel, dry wipe, or other cloth will also contribute to effective decontamination. This dry
 decontamination step can be performed by patients themselves and, along with clothing removal, should be done as early
 as possible.
- If contamination with liquid agent is suspected, patients should be decontaminated with water, ideally with a high-volume, low-pressure shower, including soap if available, gentle rubbing with a soft cloth or sponge, and active drying with a clean towel after the shower.
- If Reactive Skin Decontamination Lotion (RSDL) is available, it is recommended for spot decontamination.

TREATMENT*

- Nerve agent toxicity is the result of excessive acetylcholine, causing cholinergic crisis. Therapy focuses on treating
 the excessive secretions and bronchospasm with anticholinergic medications such as atropine with dosing titrated to
 respiratory secretions and airway resistance. Pralidoxime chloride (2-PAM CI), a specific nerve agent antidote, augments
 the primary therapy of atropine; continuous infusions may be beneficial.
- Seizures should be managed with escalating doses of benzodiazepines (midazolam, lorazepam, or diazepam).
 All patients, even without convulsions, who meet the severe criteria should be treated with midazolam, lorazepam, or diazepam 10 mg IV/IM/IO. A pediatric patient in this setting is defined as an individual less than 18 years old AND with an ideal body weight (IBW) of ≤ 40 kg. If IBW is > 40 kg, adult medication and dosing are more appropriate. For patients under 40 kg, use midazolam only: 0-13 kg -70 mcg/kg, >13-40 kg 5 mg.
- Autoinjectors (AI) are a convenient means of rapidly administering drugs to treat nerve agent exposure, which may be
 especially useful pre-hospital or at a hospital managing a large number of patients. However, only certain drugs in specific
 doses are available in autoinjectors: DuoDote or Antidote Treatment Nerve Agent Autoinjector (ATNAA) or Mark 1 kit
 (atropine 2 mg/2-PAM CI 600 mg); atropine 2 mg, 1 mg, or 0.5 mg; 2-PAM CI 600 mg; diazepam 10 mg.

EMS AGENCIES SHOULD FOLLOW THEIR ESTABLISHED TREATMENT PROTOCOLS

*National Model EMS Clinical Guidelines are also acceptable: https://www.nasemso.org/Projects/ModelEMSClinicalGuidelines/

NOTE: dosages in the model clinical guidelines are based on National Library of Medicine references; dosages noted below are based on the Agency for Toxic Substances and Disease Registry's medical management guidelines.

Patient Age	Antidotes ¹		Other Treatment
	Mild / Moderate Symptoms ²	Severe Symptoms ³	
Infant (0-2 yrs)	Atropine: 0.05 mg/kg IV/IM/IO 2-PAM CI: 15 mg/kg IV/IM/IO	Atropine: 0.1 mg/kg IV/IM/IO 2-PAM CI: 25 mg/kg IV/IM/IO	Assisted ventilation should be started after administration of antidotes for severe exposures. Repeat atropine (2 mg IV/IM) at 5- 10 minute intervals until secretions have diminished and breathing is comfortable or airway resistance has returned to near normal.
Child (2-10 yrs)	Atropine: 1 mg IV/IM/IO 2-PAM CI: 15 mg/kg IV/IM/IO	Atropine: 2 mg IV/IM/IO 2-PAM CI: 25 mg/kg IV/IM/IO	
Adolescent (>10 yrs)	Atropine: 2 mg IV/IM/IO/AI; 2-PAM CI: 15 mg/kg IV/IM/IO	Atropine: 4 mg IV/IM/IO/AI; 2-PAM CI: 25 mg/kg IV/IM/IO	
Adult	Atropine: 2 to 4 mg IV/IM/IO/AI; 2-PAM CI: 600 mg IV/IM/IO/AI	Atropine: 6 mg IV/IM/IO/AI; 2-PAM CI: 1800 mg IV/IM/IO/AI	
Elderly, Frail	Atropine: 1 mg IV/IM/IO 2-PAM CI: 10 mg/kg IV/IM/IO	Atropine: 2 to 4 mg IV/IM/IO 2-PAM CI: 25 mg/kg IV/IM/IO	

- 1 If 2-PAM CI solution is needed, prepare from a vial containing 1 g desiccated 2-PAM CI: reconstitute with 20 ml sterile water.
- 2 Mild/Moderate symptoms include localized sweating, muscle fasciculations, nausea, vomiting, weakness, dyspnea.
- 3 Severe symptoms include unconsciousness, convulsions, apnea, flaccid paralysis.

Specific Pediatric Considerations

For pediatric patients, existing autoinjectors may provide more than the recommended doses of atropine and pralidoxime. The reference below provides a strategy to mitigate this issue if time and resources allow. This method allows you to discharge the contents of autoinjectors and dilute the drug to prepare the proper dose. Expert opinion would still recommend that, given the benefit compared to the possible harm in delaying treatment, severe patients should be treated with autoinjectors even if they provide doses above recommendations.

Corvino TF, Nahata MC, Angelos MG, Tschampel MM, Morosco RS, Zerkle J, Nelson RN. Availability, stability, and sterility of pralidoxime for mass casualty use. Ann Emerg Med. 2006 Mar; 47(3):272-7.

ADDITIONAL CONSIDERATIONS

- If faced with a mass casualty incident and if pharmaceutical therapies become exhausted, consider contingency medical countermeasures at your discretion.
- Poison Control Centers provide 24-hour-a-day patient care support at 1-800-222-1222.
- The Secretary of Health and Human Services issued a declaration, effective April 11, 2017, under the Public Readiness and Emergency Preparedness Act (PREP Act) to provide liability immunity to certain individuals and entities against any claim of loss relating to the use of medical countermeasures against nerve agents, given certain conditions are met: https://www.federalregister.gov/documents/2017/05/10/2017-09455/nerve-agents-and-certain-insecticides-organophosphorous-andor-carbamate-countermeasures

OTHER RESOURCES

U.S. Department of Health and Human Services

Centers for Disease Control and Prevention / Agency for Toxic Substances and Disease Registry

https://emergency.cdc.gov/agent/nerve/index.asp

https://www.atsdr.cdc.gov/mmg/mmg.asp?id=523&tid=93

Office of the Assistant Secretary for Preparedness and Response

https://chemm.nlm.nih.gov/na_hospital_mmg.htm

Personal Protective Equipment

U.S. Department of Labor

Occupational Safety and Health Administration

https://www.osha.gov/Publications/OSHA3370-protecting-EMS-respondersSM.pdf

https://www.osha.gov/pls/oshaweb/owadisp.show_document?p_table=standards&p_id=9765

Patient Decontamination

U.S. Departments of Health and Human Services and Homeland Security

https://www.phe.gov/Preparedness/responders/Pages/patientdecon.aspx

General

U.S. Department of Health and Human Services

Office of the Assistant Secretary for Preparedness and Response

https://asprtracie.hhs.gov/

National Fire Protection Association

https://www.nfpa.org/codes-and-standards/all-codes-and-standards/list-of-codes-and-standards/detail?code=473