

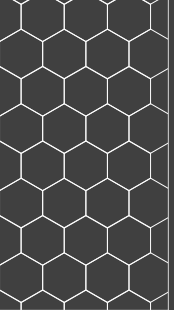


EVOLVING OVERDOSE RESPONSE

ADAPTING TO SEDATIVES IN THE DRUG SUPPLY

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Rose Laurano, MPH, Harm Reduction Manager, Division of Substance Use Prevention and Harm Reduction, Philadelphia Department of Public Health



Learning Objectives

1. Understand the effects of sedatives on overdose presentation and recognition.
2. Evaluate the effects and limitations of naloxone in sedative-involved overdoses.
3. Identify gaps in current opioid overdose training and provide guidelines for responding to sedative-involved overdoses.

Today's training was adapted from the Philadelphia Department of Public Health.

Today's Presenters



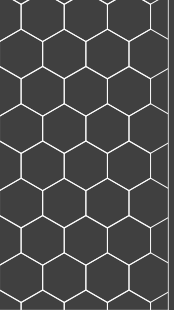
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Community-informed approaches to the escalating presence of xylazine in opioids

NIDA-funded study (R21DA058582)

Thomas Jefferson University (PI Reed) | Brandeis University (Co-I Green)

Step 1: Interviewed people using xylazine and other content experts in Philadelphia, PA, a xylazine-saturated market.

Step 2: Presented results to people using xylazine and other content experts. Used co-design to develop three interventions.

Step 3: Piloted interventions in urban, suburban, and rural communities with emerging xylazine markets in Massachusetts



1

XYLAZINE & OTHER SEDATIVES

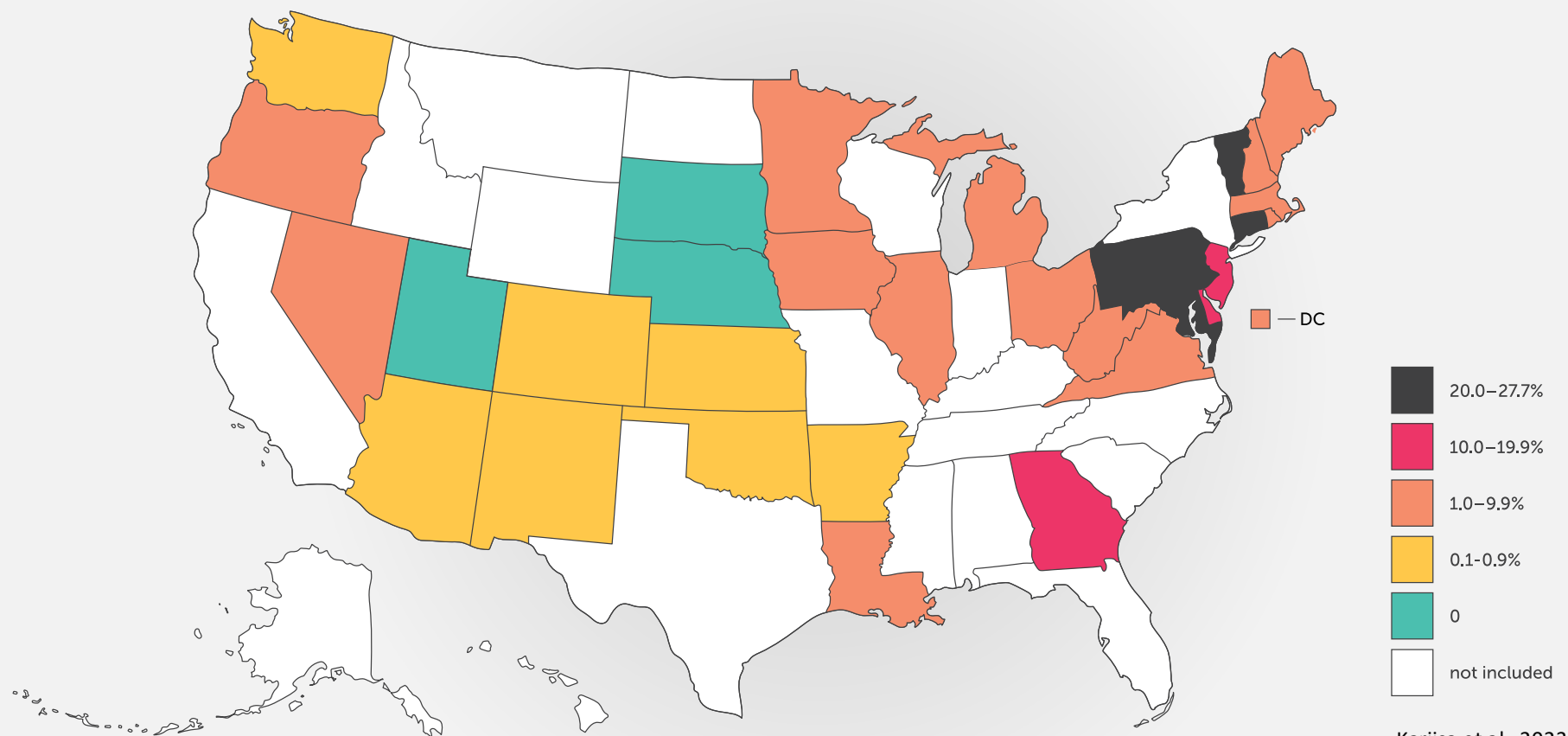
Overdose crisis in Philadelphia and the changing drug supply

TIMELINE



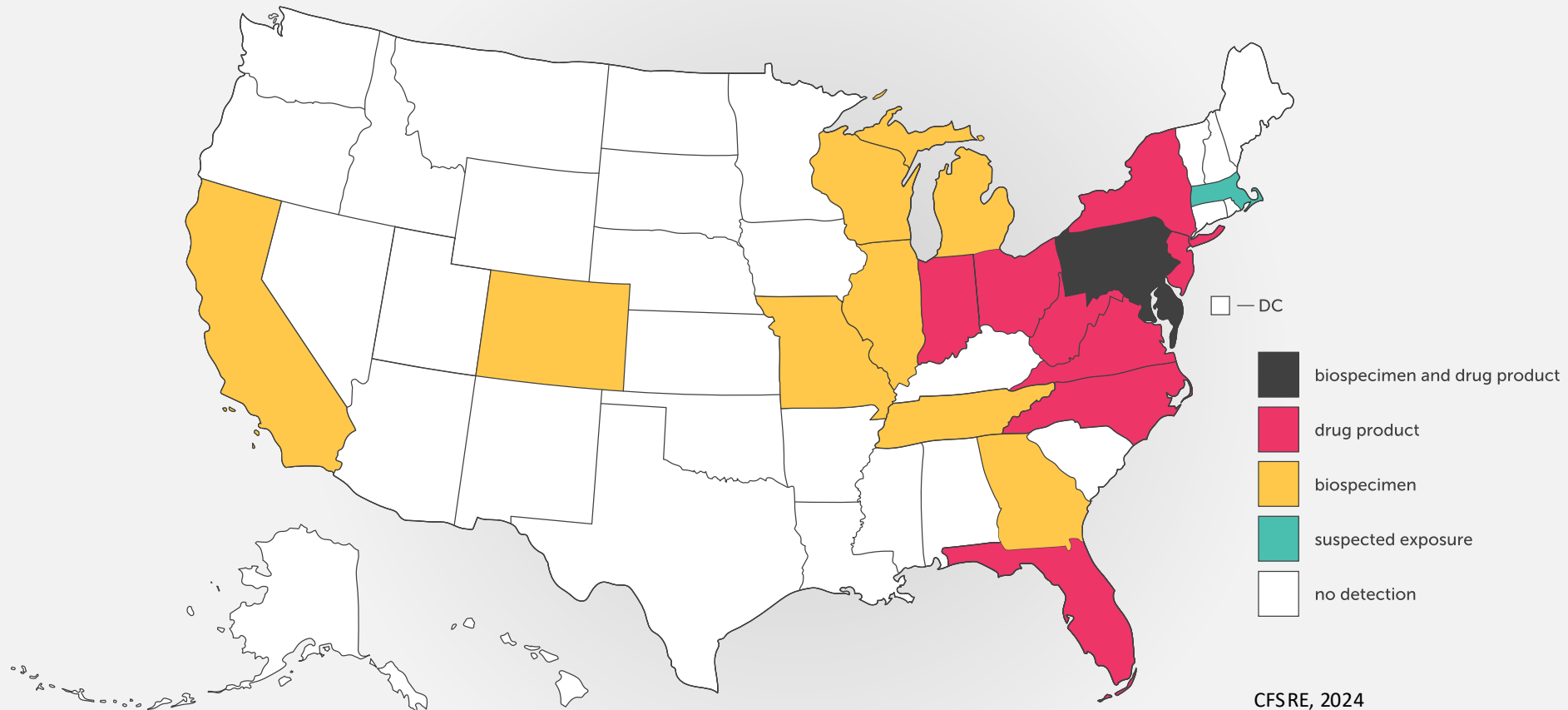
Xylazine's spread across the US, January 2019–June 2022

PERCENTAGE OF IMF-INVOLVED OVERDOSE DEATHS WITH XYLAZINE DETECTED



Medetomidine's spread across the US, late 2022–May 2024

GEOGRAPHICAL DISTRIBUTION OF MEDETOMIDINE EMERGENCE



What is dope in most places?

SYNTHETIC OPIOID

fentanyl
nitazenes
carfentanil

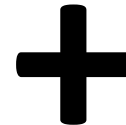
DOES respond
to naloxone



SEDATIVE

xylazine
medetomidine
benzodiazepines

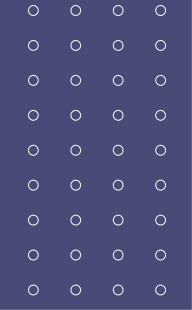
DOES NOT respond
to naloxone



ADULTERANT

benzocaine
caffeine
lidocaine
quinine

DOES NOT respond
to naloxone



Xylazine and Medetomidine

THE SAME...

- Veterinary medication
- Alpha-2-agonist
- Not responsive to naloxone
- Rapid onset
- Causes prolonged sedation
- Almost always found with fentanyl
- Involved in fatal overdose
- Causes withdrawal
- Not federally scheduled

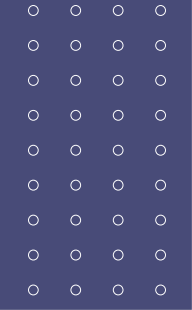
BUT DIFFERENT

XYLAZINE

- Clear association with wounds
- Scheduled in PA

MEDETOMIDINE

- 100-200x more potent than xylazine
- Longer sedation
- More pronounced lower heart rate
- Anecdotal reports of association with hallucinations and worse withdrawal symptoms



How do xylazine and medetomidine affect overdose?

Cause prolonged sedation

People need positioning to maintain open airway

Sedatives are not responsive to naloxone

Response needs to place great emphasis on breathing

People may remain non-responsive

2

ADAPTING OVERDOSE RESPONSE

Reminder: responding to an overdose

1. Call out for help.
2. Administer naloxone.
3. Call 911.
4. Start rescue breathing.
5. After three minutes if not breathing, give 2nd dose of naloxone.
6. Continue rescue breathing until breathing is restored.
7. Stay with them until help arrives.
8. Provide a compassionate comeback.

Sedative-involved overdose recognition



FOCUS ON RESPONSE

SYNTHETIC OPIOID OVERDOSE

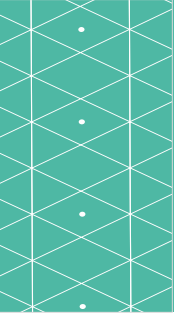
1. Check for responsiveness
2. Check for breathing
3. Administer naloxone
4. Call 911
5. Administer rescue breathing



FOCUS ON BREATHING

SYNTHETIC OPIOID OVERDOSE + SEDATIVE OVERDOSE

1. Check for breathing
2. Check for responsiveness
3. Administer naloxone
4. Call 911
5. Administer rescue breathing



Recognizing the emergency

The emergency is that they are not breathing,
not that they aren't answering you.

Change your approach.

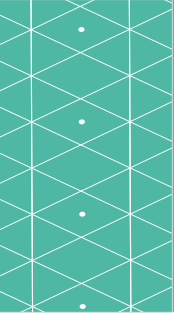
**Focus on respiration
instead of responsiveness.**

Methods to assess breathing

LOOK. Watch for breathing. Look for chest movement OR watch shoulders (rise & fall). Count to make sure it's at least 1 breath every 5 seconds.

LISTEN. Can you hear them breathing? Does it sound strained or choked?

CHECK. Hold a piece of glass (like a small mirror), or metal object (like a key), or piece of plastic (rescue breathing mask)—or any other item that collects condensation—in front of their mouth.



Guidelines for responding to sedative-involved overdoses

**Don't just ask if they're okay.
Check to see if they're blue or gray.**

Train responders to look for color changes and respiration.

**Confirm that at least 1 breath is taken every 5 seconds.
Check "lips & tips" for blue or gray.**

Gaps in current standard overdose response training

The priority must shift from eliciting a response to assessing, and supporting, breathing/oxygenation.

Assessing for responsiveness with painful stimuli can cause unnecessary harm.

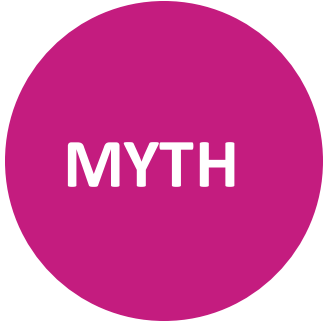
Responsiveness is no longer the goal of overdose response.

There IS such a thing as too much naloxone.



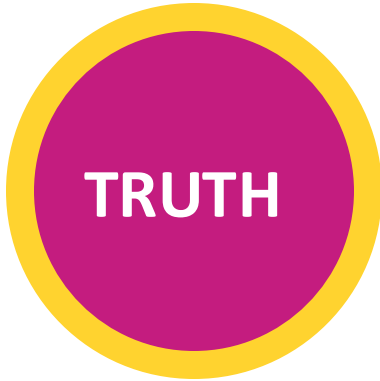
Photo credit: The Everywhere Project

Naloxone works!



MYTH

Naloxone doesn't work on sedative-involved overdoses.



TRUTH

Naloxone is working to restore breathing from the opioid overdose **AND** the person may still be sedated on xylazine or another sedative.

Naloxone will not resolve sedation, but it will help people breathe during overdose situations.

Naloxone works, but... make a plan for monitoring to keep the person safe

Naloxone takes time to work and should be supported by rescue breathing.

Naloxone wears off in 45 min–2 hours, but the person may still be sedated.

Sedation resolves with time, which may be up to 2–3 hours, possibly more with medetomidine.

PRO TIP: have a plan ready for monitoring or transition.

Too much naloxone can be harmful

A 2023 study comparing 4 mg to 8 mg of an initial intranasal (IN) dose found that people who received the higher dose were 2.5 times more likely to experience withdrawal symptoms, with no difference in survival. (Payne et al., 2024)

The ideal dose of naloxone is one that restores breathing without inducing withdrawal. (Russell et al., 2024)

Naloxone is totally safe for people who are not physically dependent on opioids, but over-administration can cause precipitated withdrawal among PWUO.

Too much naloxone can be harmful

The more naloxone administered, the worse the withdrawal.

Studies have shown that people who experience severe precipitated withdrawal are more at risk for fatal overdose:

- People may try to immediately use more to relieve their symptoms, causing them to overdose again when the naloxone wears off.
- People may try to use in secluded places/away from people out of a concern that they will receive too much naloxone.

**Don't be afraid to give someone naloxone if they are not breathing!
But try and use as little as possible.**

What is Compassionate Comeback?

When responding, administer the least amount of naloxone possible.

Be kind

Talk people through the process, even if they can't verbally respond

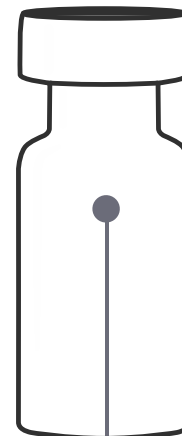
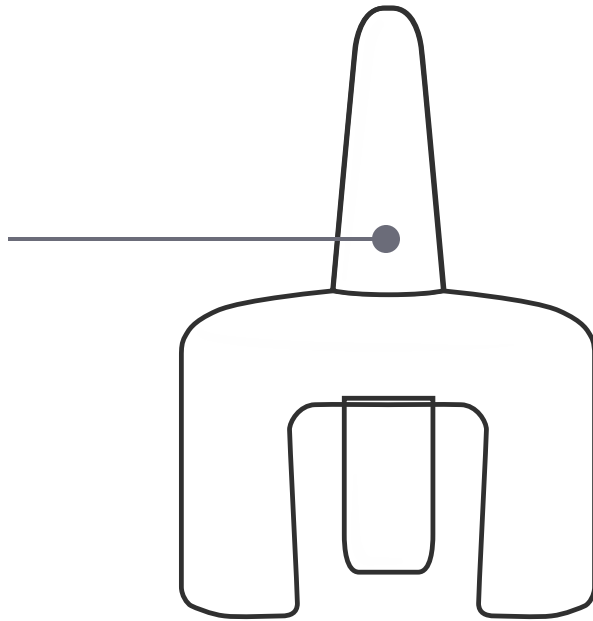
Supportive aftercare:

- Create a calm environment to the extent possible.
- Reduce the number of people nearby, especially those in uniform.
- Give them space and welcome them back.
- Explain what happened.
- Respect their wishes—don't make them stay.
- Take care of yourself and your needs.

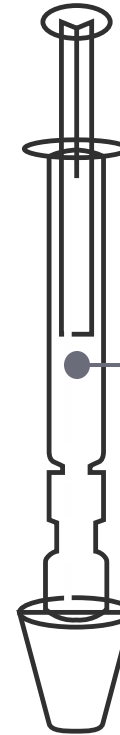
Different forms of naloxone—Best

3mg

nasal
spray
(brand)



0.4mg intramuscular
(IM) (generic)

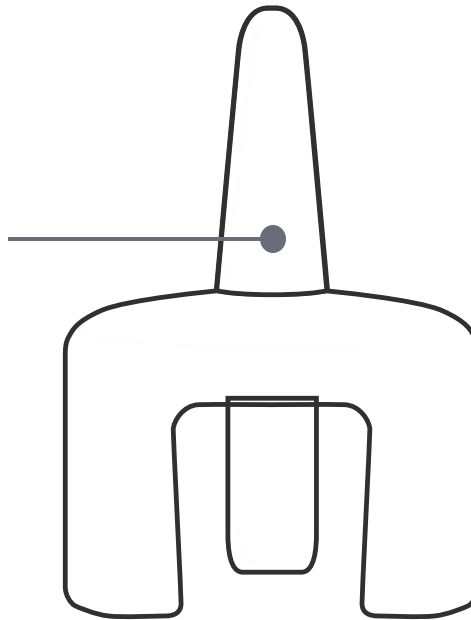


1mg & 2mg
nasal atomizer
(generic)

Different forms of naloxone—Good

4mg

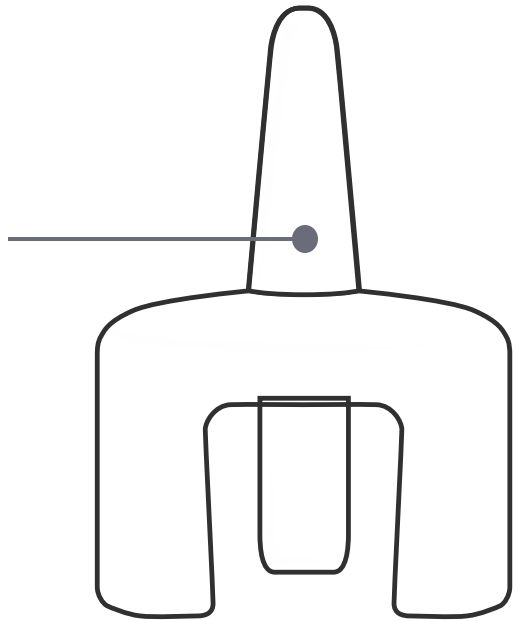
nasal spray
(brand or
generic)



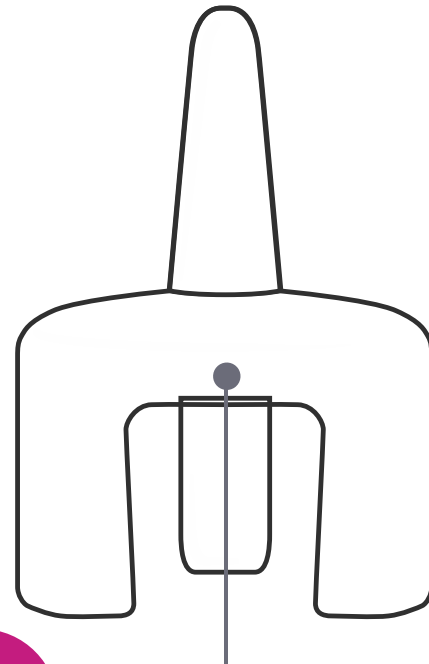
Different forms of naloxone —

Not
recommended

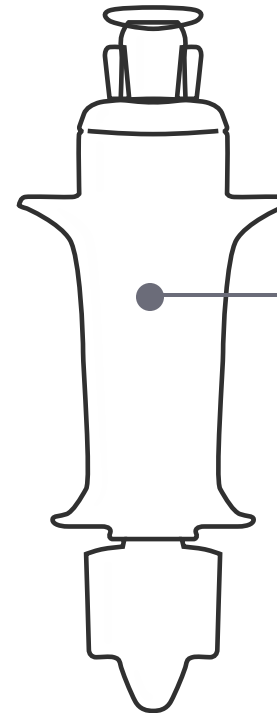
nalmafene



8mg



nasal applicator
(brand)

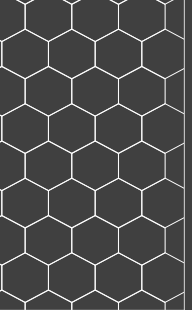


5mg auto
injector
intramuscular
(IM) (brand)



3

EXPANDING YOUR TOOLBOX



Look at your toolbox: two scenarios

MY TOOLBOX IS BASIC. I CAN'T CHANGE TOO MUCH.

Budget limitations

Training limitations

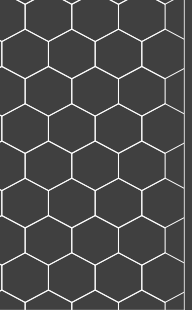
Capacity limitations

WE'RE LOOKING TO EXPAND OUR OPTIONS!

Available funding

Training capabilities

Invested stakeholders



Limited in changing your standard kit?

STANDARD KIT

Barrier/CPR mask

Naloxone

Gloves

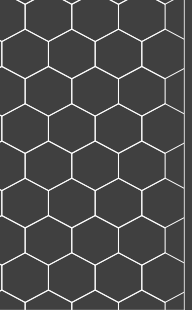
WHAT TO ADD:

Emphasis on breathing support and assessment

Patience! Let the naloxone work

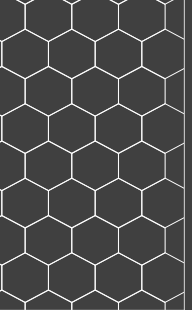
Positioning (head tilt, rescue position)

Methods to reduce the harms of sedation



Methods to reduce the harms of sedation

1. If laying, utilize the recovery position.
2. If sitting, head reposition (decompress windpipe).
3. Check that limbs aren't "pinned".
4. Massage limbs during prolonged sedation.
5. Move them to a safe place (e.g., away from/out of street, away from train tracks, not in the middle of the sidewalk).
6. If possible, ask someone to keep an eye on them.



Comfortable in the grey zone

More uncertainty than there used to be—get comfortable there.

The person was not overdosing but is severely sedated.

You can still reduce the harm!

After reversing an overdose:

Naloxone can wear off and the person is still sedated.

Have a couple of ready-to-go transition plans.

Can you get them to a community hub/nearby resource?

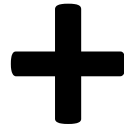
Are there any bystanders you can ask to check on them?

+/- Call 911/EMS

Can you find a safe place for them to sit?

Comfortable in the grey zone

YOU DO THE BEST
YOU CAN FOR THE
SITUATION YOU
ARE IN



AND THAT
DOESN'T ALWAYS
MEAN MORE
NALOXONE

Able to adapt your standard kit?

STANDARD KIT

Barrier/CPR mask

Naloxone

Gloves

ADDITIONAL TOOLS

Pulse oximeter

Ambu-bag

Oropharyngeal & nasopharyngeal airways*

Recreational oxygen cans

Oxygen tanks*

** These tools require extensive and specialized training. Talk to a clinician about the best way to access this.*

Pulse oximeters

PROS

Provides ongoing data to refer to during overdose response

Can help assess oxygen levels if you're unsure

Assists with monitoring breathing when in a gray zone

CONS

Requires training to use

Has higher supply costs

Unreliable

Skin color

Nail polish

Injury/Callouses



AMBU Bag

PROS

Better respiration
Alternative for those who won't do rescue breathing

CONS

Training: seal and frequency
One vs. two people
Single use
Expensive (\$15-\$30)



Oropharyngeal & nasopharyngeal airways

PROS

Keeps airways free of obstruction

Facilitates ventilation (works well with masks and bags)

Quick to insert

Cost effective (\$1-\$10)

CONS

Can trigger a gag reflex

Specialized training/expertise required

Incorrect placement can cause trauma/obstruct airway

Not suitable for all people



Recreational oxygen cans

PROS

Uses beyond opioid overdose response

More accessible and affordable than oxygen tanks

Less bulky than oxygen tanks

Good complement to rescue breathing and airway management

CONS

Supplemental oxygen only

Bulky for unhoused people

Should not be used near open flame

Moderately expensive for widespread distribution (\$18-\$20)



Oxygen tanks

PROS

Best respiration

Alternative for those who won't do rescue breathing*

Reduces the need for naloxone

**Only for people with depressed respiration—i.e., breathing but not enough. If someone is not breathing, this will not help.*

CONS

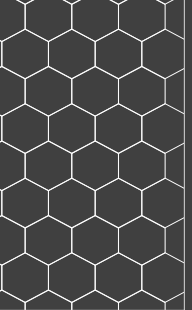
Specialized training

Potential dangers (flammable)

Difficult to transport

Very expensive (\$400-\$4,000)





What to remember...

Naloxone works!

Breathing > responsiveness.

Sedation is not the emergency.

The grey zone is bigger.

Naloxone dose matters (there is such a thing as too much).

Remember: compassionate comeback

More tools doesn't mean better tools.



4

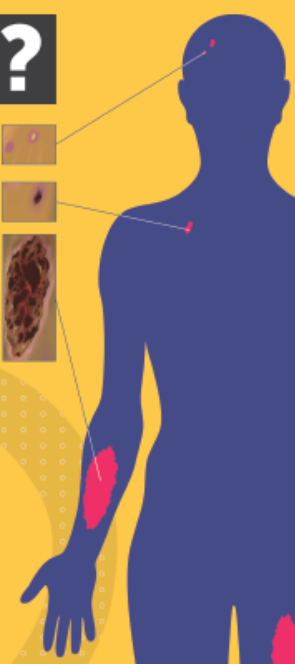
OTHER XYLAZINE INTERVENTIONS

Daily Skin Check poster

HAVE YOU CHECKED YOUR SKIN TODAY?

XYLAZINE CAUSES WOUNDS
and is found in the drug supply.
Xylazine wounds can form
anywhere on your body and
usually start as a small spot.
Catch wounds early and talk
to a healthcare provider or
your local harm reduction
organization for more
information.

STAGE 1
STAGE 2
STAGE 3



To learn more:
[Jefferson.edu/XylazineResponse](https://jefferson.edu/XylazineResponse)

English Español

Thomas Jefferson University

¿HAS REVISADO TU PIEL HOY?

LA XILACINA GENERA HERIDAS
Y se encuentra actualmente
en el mercado de drogas. Las
heridas causadas por xilacina se
pueden generar en cualquier
parte de tu cuerpo, y usualmente
empiezan como una mancha
pequeña. Detecte las heridas
tempranamente, y pónganse en
contacto con un profesional de
la salud o su organización de
reducción de daño local.

ETAPA 1
ETAPA 2
ETAPA 3



Para aprender más:
[Jefferson.edu/SobreXilacina](https://jefferson.edu/SobreXilacina)

English Español

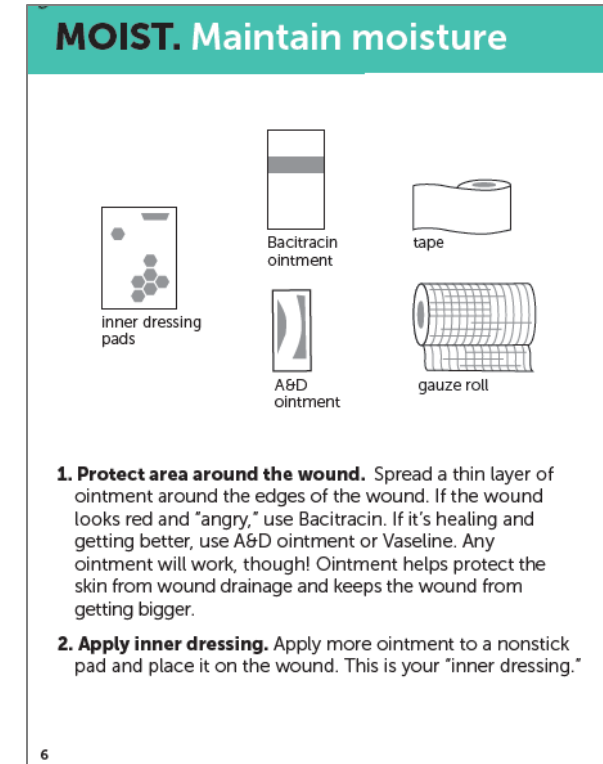
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Wound Care booklet

BACK/FRONT COVERS



INSIDE



Wound Care video



STEP 1

Preparing for Treatment

Downloadable materials

ALL MATERIALS ARE FREE AND AVAILABLE AT:

www.Jefferson.edu/xylazineresponse



www.Jefferson.edu/sobrexilacina



ACKNOWLEDGEMENTS

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Co-design participants

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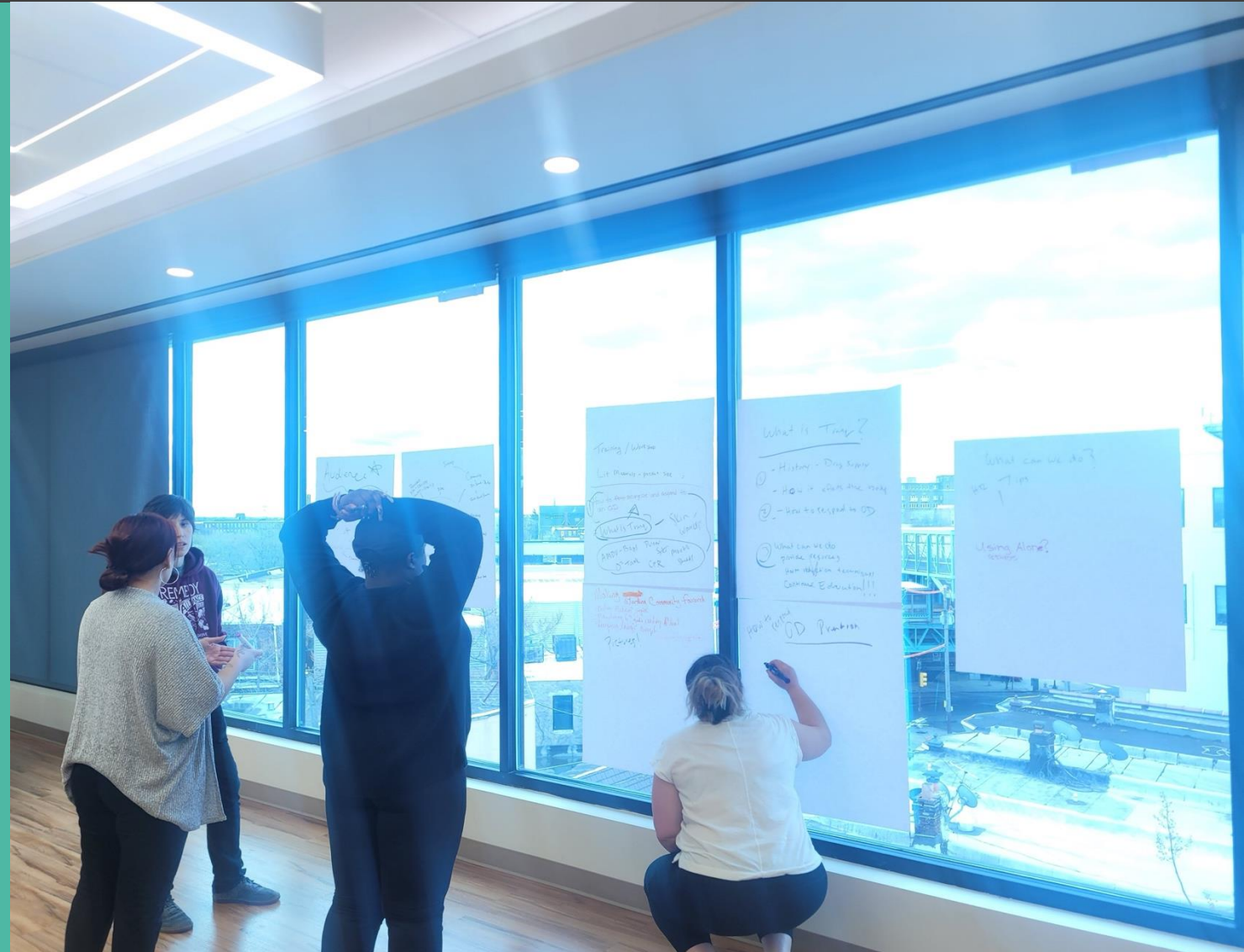
Berkshire Harm Reduction

Fenway Health

Healthy Streets

Tapestry

National Institute on Drug
Abuse: (R21DA058582)



QUESTIONS?

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THANK

YOU!