

#### Keeping Current With Ever-Changing NPS Landscapes – Evaluating Trends and Connecting Case Histories

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#### Disclosure

• I have nothing to disclose.



#### Background



### **Our NPS Discovery Program**

#### **Forensic Toxicology Investigations**



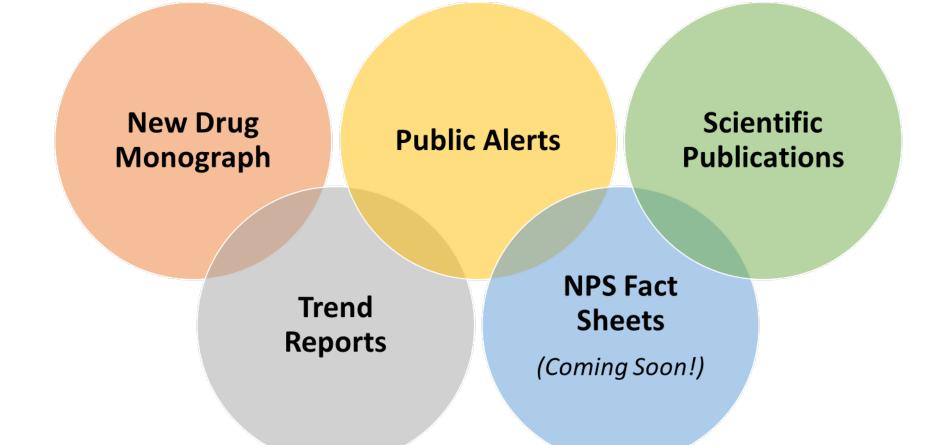




- 1. Intelligence
  - What is out there?
- 2. Surveillance
  - Have we seen it?
- 3. Monitoring
  - How often?
- 4. Response
  - What do we do?
- 5. Forecasting
  - What's next?

Cfsre O NPS DISCOVERY







#### Methods



# Methods

- Identification
  - SCIEX TripleTOF<sup>®</sup> 5600+
    LC-QTOF-MS
  - >875 drug panel



#### • Quantitation

- Waters Xevo TQ-S micro LC-MS/MS
- Standard addition





# **Toxicology Workflow**

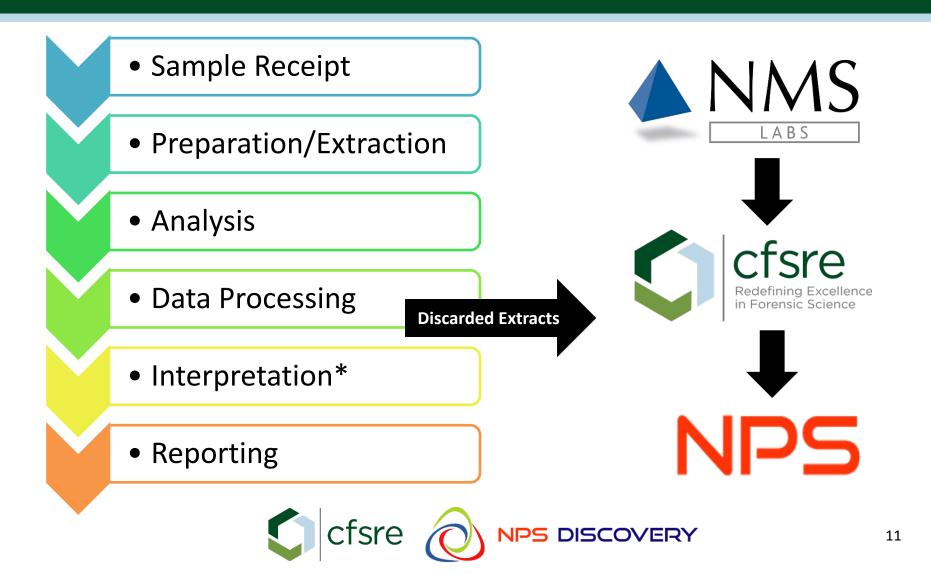
- Sample Receipt
- Preparation/Extraction
- Analysis
- Data Processing
- Interpretation\*
- Reporting



#### **Sample Sources**

- NMS Labs
- Medical Examiner
- Coroner
- Hospitals
- Poison Centers
- Other\*

# **Toxicology Workflow**



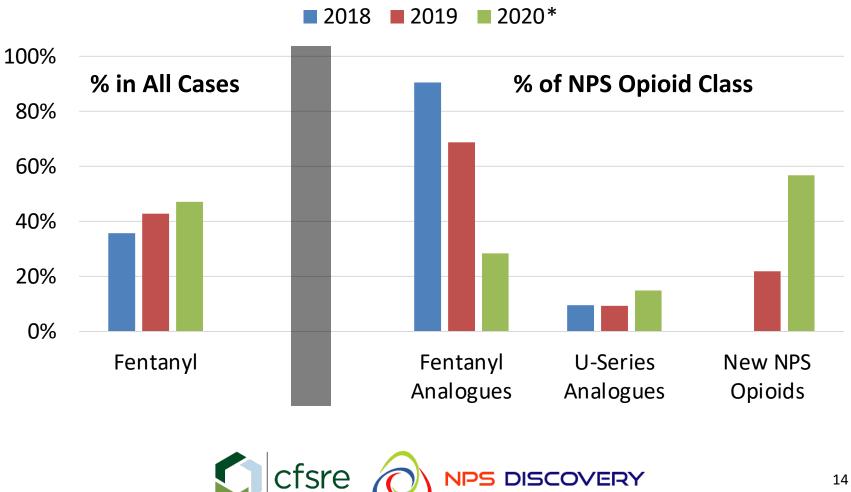
#### NPS Trends



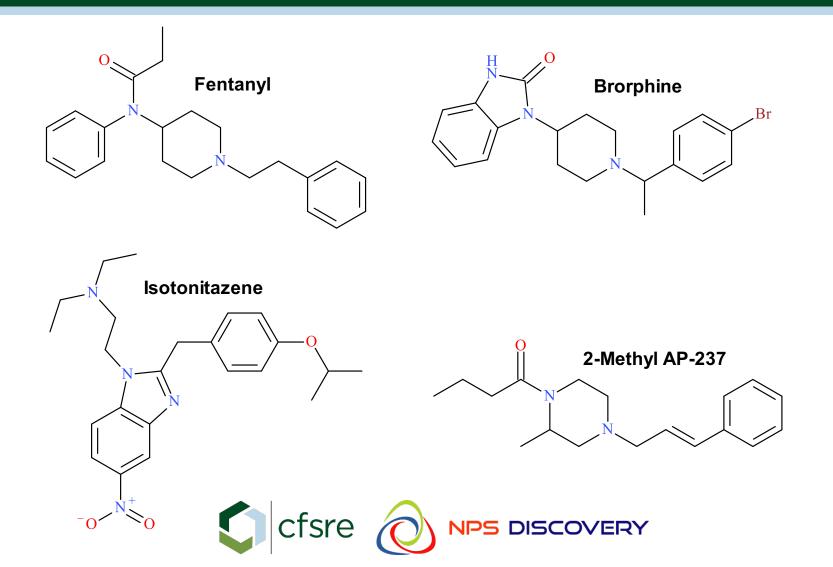
### NPS Benzodiazepines

2018	2019	2020*	*Only 8 months
			% of class
📘 Etizolam (47%)	Etizolam (56%)	🔲 Etizolam (45%) 🗸	
🔲 Flubromazolam (26%)	🔲 Flualprazolam (13%) 个	🔲 Flualprazolam (40%) 个	
🔲 Diclazepam (11%)	$\square$ Flubromazolam (12%) $oldsymbol{\downarrow}$	🔲 Flubromazolam (7%) 🗸	
🔲 Clonazolam (7%)	🔲 Bromazolam (9%) 个	🔲 Clonazolam (3%) 个	
🔲 Flubromazepam (5%)	🔲 Diclazepam (6%)	🔲 Adinazolam (2%) 个	
🔲 Phenazepam (3%)	🔲 Clonazolam (2%)	Pyrazolam (1%)	
🔲 Pyrazolam (1%)	🔲 Flubromazepam (1%)	🔲 Bromazolam (1%)	
🔲 Flualprazolam (1%)	🔲 Phenazepam (1%)	🔲 Phenazepam (1%)	
		Estazolam (>1%)	
		🔲 Metizolam (>1%)	
		Diclazepam (>1%)	
		🔲 Flubromazepam (>1%)	

# **NPS** Opioids



### NPS Opioids



# Isotonitazene $\rightarrow$ Brorphine

- More information on isotonitazene cases:
  - Session 6: Analytical Toxicology
  - Wednesday, September 23<sup>rd</sup>
  - 10:45 11:00am PT (1:45PM 2:00PM ET)
  - Isotonitazene: A Novel Benzimidazole μ-Opioid Agonist as the Latest Novel Opioid
  - Barry K. Logan, Donna M. Papsun, Alex J. Krotulski, Sherri L. Kacinko

Article

#### Isotonitazene Quantitation and Metabolite Discovery in Authentic Forensic Casework

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# Isotonitazene $\rightarrow$ Brorphine

- "Linking case histories"
- Isotonitazene first emerged in the Midwest in mid-to-late 2019
- Received a powder to test
  - Positive for isotonitazene, flualprazolam, and heroin  $\rightarrow$
- An "autopsy" of the cases
  - 89% of samples were positive for isotonitazene + flualp./etizolam





# Isotonitazene $\rightarrow$ Brorphine

- Fast-forward to mid 2020
- DEA North Central Lab reports a gray powder (sold as heroin/fentanyl) containing flualprazolam and brorphine
- Drug forums began suggesting that manufacturers are moving away from isotonitazene
- Brorphine has become the replacement for isotonitazene
   To date: ~40 cases

make this logistically if not impossible at least tremendously difficult. As such intramuscular tend to be my route of choice. That said I now find myself at something of an impasse. The supplier from whom I have been procuring my isotonitazene has run out and has no plans to restock. As such, I am in search of a replacement what are the similar range of potency. I have tried 2-methyl-AP-237 and while I

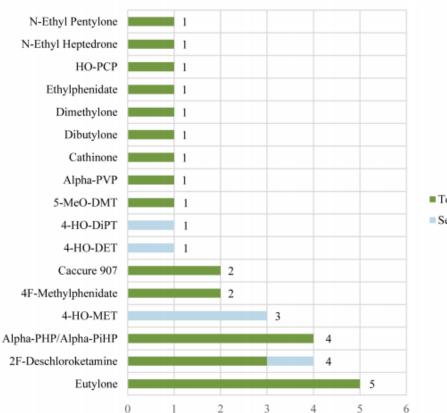


# 2-Methyl AP-237

- To date, three cases with histories
- Death investigation in California
  - 29 y/o M; Discovered deceased in sober living facility
  - Blood concentration: 5,800 ng/mL (No other DoA or NPS)
- Clinical non-fatal overdose scenario in Georgia
  - Ingested 1g of 2MAP; Presented to ED; Administered naloxone, treated, and released; Returned with different drug overdose
  - Blood concentrations: 2.9 to 35 ng/mL
- Death investigation in Tennessee
  - 35 y/o M; Hx drug use; Drove car into ditch; Found deceased in home
  - Blood concentration: 1,100 ng/mL
  - 3-HO-PCP (11 ng/mL) and 2F-Deschloroketamine (3.9 ng/mL)



# NPS Stimulants / Hallucinogens



#### NPS Stimulant & Hallucinogen Positivity

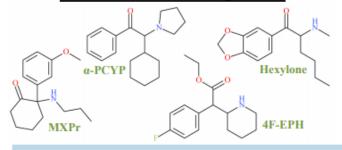
Toxicology

#### Seized Drug

#### NPS Stimulant Combinations

Combination	Frequency
Eutylone + Etizolam + Fentanyl + Cocaine	1
Eutylone + Etizolam + Cocaine	1
Eutylone + Fentanyl + Cocaine	1
Eutylone + Fentanyl	1

#### New Discoveries in Q2 2020



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# Synthetic Cannabinoids

2018	2019	2020*	
<b>5F-ADB (52%)</b>	5F-MDMB-PICA (52%)	5F-MDMB-PICA (55%)	
5F-MDMB-PICA (15%)	4F-MDMB-BINACA (31%)	🔲 4F-MDMB-BINACA (31%)	
MMB-FUBINACA (13%)	FUB-AMB (8%)	🔲 MDMB-4en-PINACA (10%)	
🔲 ADB-FUBINACA (7%)	🔲 APP-BINACA (3%)	🔲 4F-MDMB-BICA (1%)	
<b>5F-EDMB-PINACA (2%)</b>	MDMB-4en-PINACA (2%)	🔲 ADB-FUBINACA (1%)	
	🔲 ACHMINACA (1%)		
*Only 8 months % of class Not comprehensive lists			

#### Conclusions



# Conclusions

- The NPS landscape in the U.S. remains dynamic and fluid
- NPS Benzodiazepines
  - Expansion and diversification of the class
  - Primary drug dominates market
- NPS Opioids
  - Fentanyl analogues are gone; New opioids are here
  - Rate of turnover appears to be 6-9 months or less
- NPS Stimulants / Hallucinogens
  - Comparatively lower prevalence  $\rightarrow$  May depend on sample population
- Synthetic Cannabinoids
  - Primary drug dominates market; International differences
  - Turnover more common among second tier compounds





### **Questions**?

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