

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



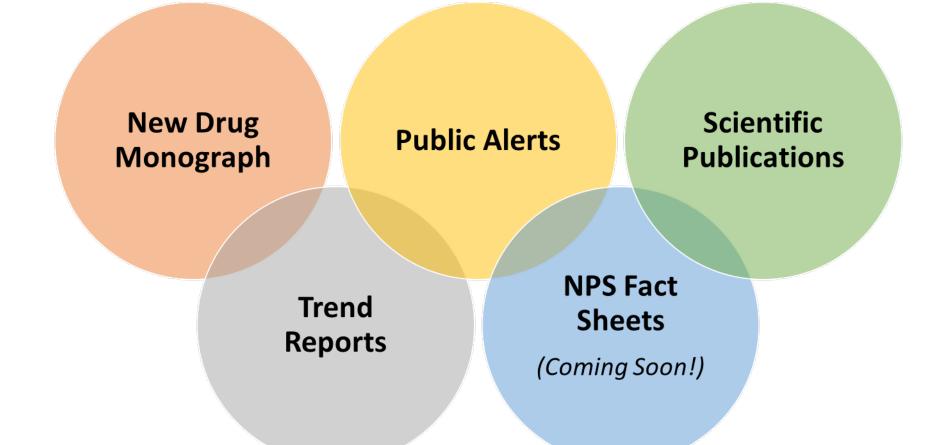




- 1. Intelligence
 - What is out there?
- 2. Surveillance
 - Have we seen it?
- 3. Monitoring
 - How often?
- 4. Response
 - What do we do?
- 5. Forecasting
 - What's next?

Cfsre O NPS DISCOVERY







Methods



Methods

- Identification
 - SCIEX TripleTOF[®] 5600+
 LC-QTOF-MS
 - >875 drug panel



• Quantitation

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





Toxicology Workflow

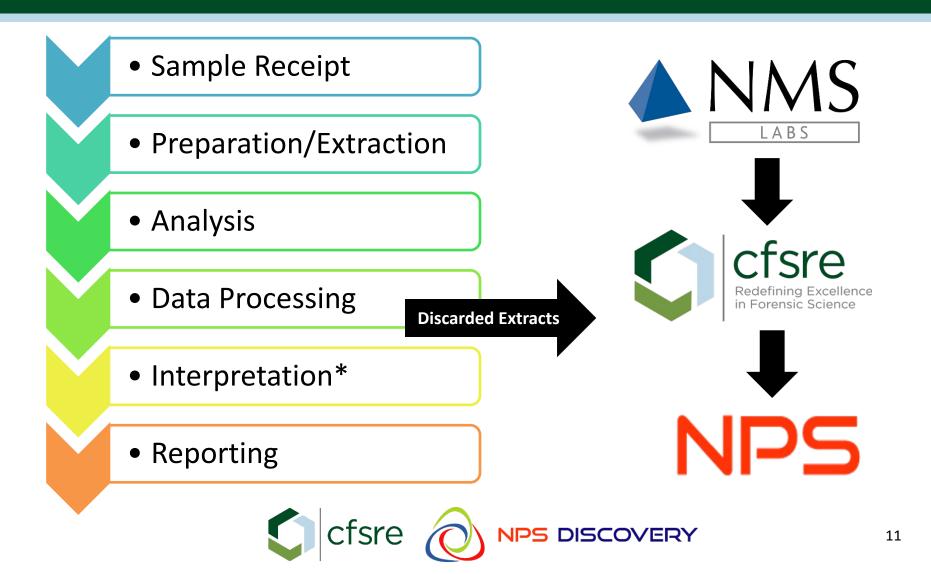
- Sample Receipt
- Preparation/Extraction
- Analysis
- Data Processing
- Interpretation*
- Reporting



Sample Sources

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

Toxicology Workflow



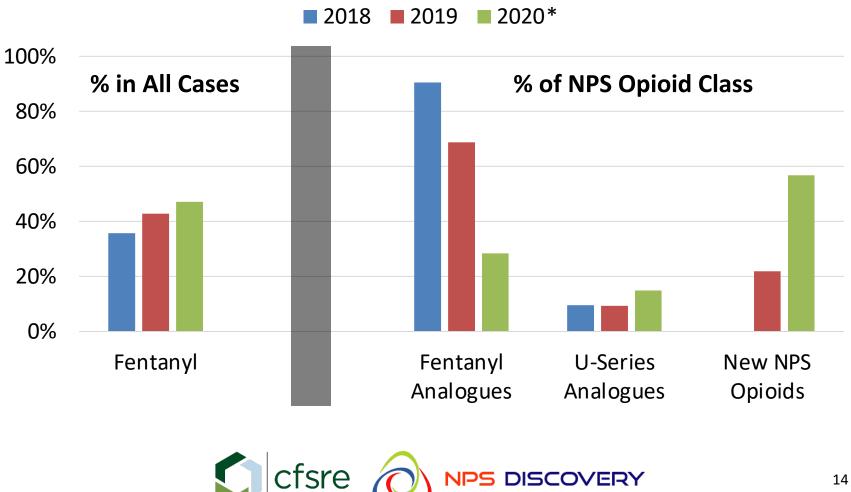
NPS Trends



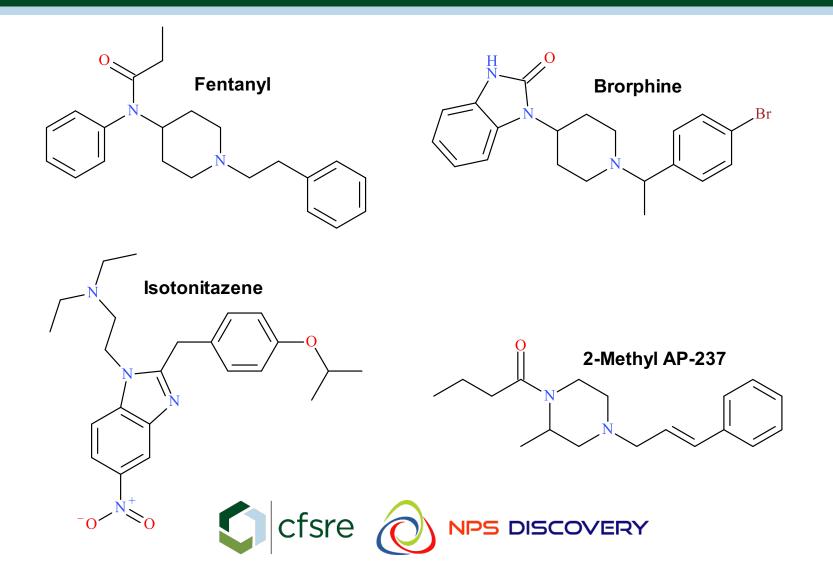
NPS Benzodiazepines

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 23rd
 - 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

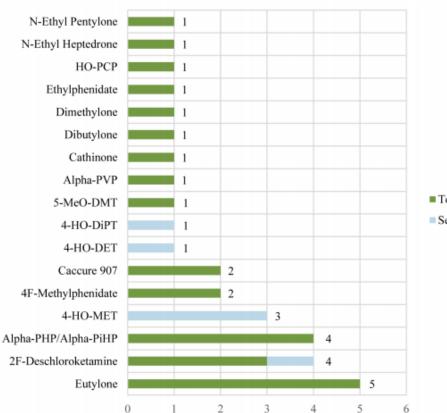


2-Methyl AP-237

- 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

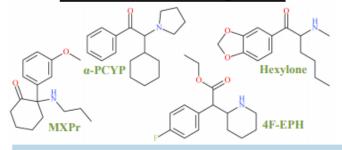
Toxicology

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*	
5F-ADB (52%)	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%)	
5F-EDMB-PINACA (2%)	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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