

## **GENERAL DISCLOSURES**

The University of Washington School of Medicine also gratefully acknowledges receipt of educational grant support for this activity from the Washington State Legislature through the Safety-Net Hospital Assessment, working to expand access to psychiatric services throughout Washington State.



## **CONFLICTS OF INTEREST?**

NONE!



## **OBJECTIVES**

- Review some common drugs of abuse popular among young adults
- Introduce some "newer" drugs of abuse
- Understand psychiatric manifestations of these substances of abuse
- Learn resources to use in order to stay current with evolving drug trends and newly synthesized drugs of abuse
- Questions, discussion



## **DESIGNER DRUG**

 Synthetic structural or function analogs of controlled substances designed to mimic the pharmocological effects of the original drug while avoiding illegality

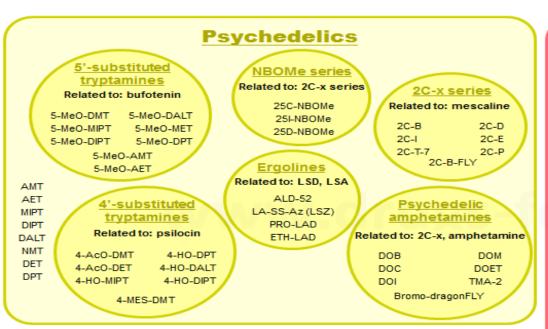


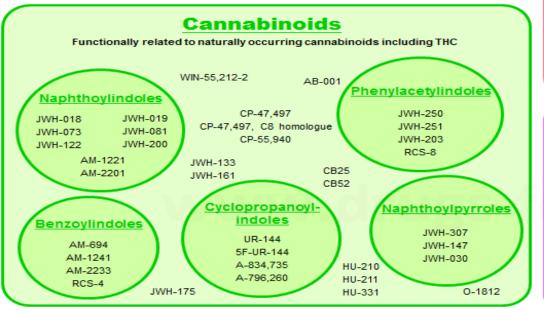
## **REASONS FOR USE**

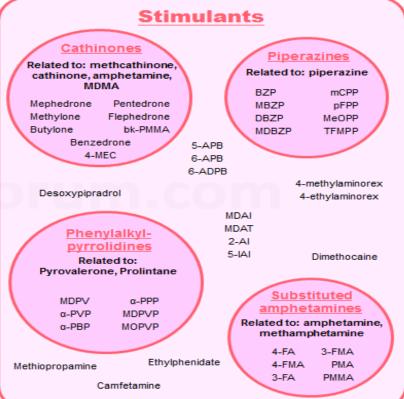
- Often Legal
- Often Less expensive
- More readily available
- More desirable pharmacological effects



## **CHALLENGES TO THIS TOPIC**







#### Dissociatives

Related to: ketamine, PCP

Methoxetamine 3-MeO-PCP 4-MeO-PCP 3-MeO-PCE 2-MeO-ketamine

# Copioids Related to: morphine, fentanyl, heroin α-methylfentanyl 3-methylfentanyl MPPP O-desmethyltramadol 7-acetoxymitragynine Metonitazene AH-7921 Phenazepam

## **SUBSTANCES COVERED TODAY**

- Synthetic Marijuana (Spice)
- Synthetic Cathinones (Bath Salts)
- MDMA (Ecstasy)
- Others
  - Piperazines
  - 2C series
  - DXM (Dextromethorphan)
  - "Syrup"
  - GHB
  - Kratom



## HOW DO I ASK MY PATIENTS ABOUT SUBSTANCES?

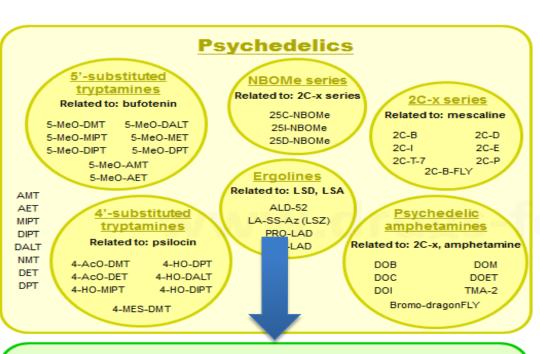
 Do you use any substance to change the way you feel or enhance your experiences?

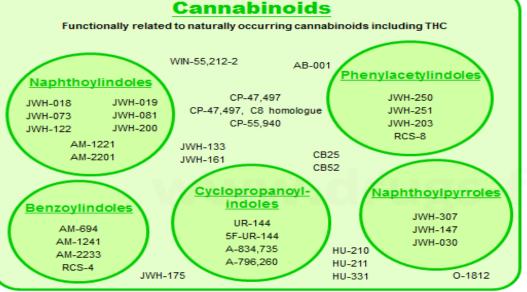
What is it? What does it do for you?

What's the newest thing?









#### **Stimulants** Cathinones **Piperazines** Related to: methcathinone Related to: piperazine cathinone, amphetamine, BZP mCPP. Mephedrone Pentedrone MBZP pFPP Methylone Flephedrone DBZP MeOPP Butylone bk-PMMA MDBZP TEMPP Benzedrone 5-APB 4-MEC 6-APB 6-ADPB 4-methylaminorex Desoxypipradrol 4-ethylaminorex MDAI MDAT Phenylalkyl-2-AI pyrrolidines 5-IAI Dimethocaine Related to: Pyrovalerone, Prolintane Substituted MDPV α-PPP amphetamines α-PVP MDPVP Related to: amphetamine. α-PBP MOPVP methamphetamine 4-FA 3-FMA Ethylphenidate 4-FMA **PMA** Methiopropamine 3-FA **PMMA** Camfetamine

#### **Dissociatives**

Related to: ketamine, PCP

Methoxetamine 3-MeO-PCP 4-MeO-PCP

3-MeO-PCE 2-MeO-ketamine

## Opioids Related to: morphine,

fentanyl, heroin

α-methylfentanyl

3-methylfentanyl

MPPP

O-desmethyltramadol

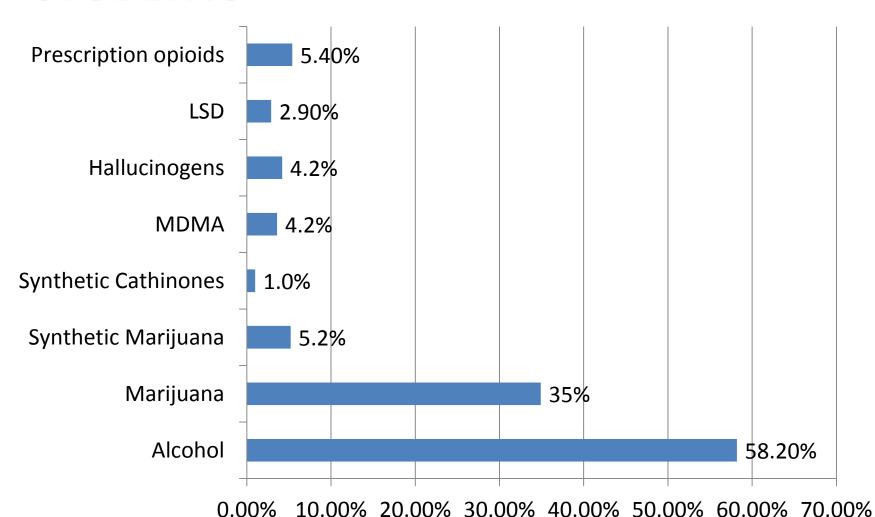
7-acetoxymitragynine

Metonitazene

AH-7921

Phenazepam

# PAST YEAR DRUG USE BY 12<sup>TH</sup> GRADE STUDENTS





## **SPICE = SYNTHETIC CANNABINOIDS**

- Herbal mixtures sprayed with chemical additives
  - "Spice"
  - "K2"
  - "Yucatan Fire"
  - "Skunk"
  - "Moon Rocks"
  - "Crazy Clown"
  - "Herbal Madness"
- Sold as
  - Potpourri
  - Incense
  - Air Fresheners
  - Obtained online, head shops, or gas stations/convenience stores
- Route of administration similar to marijuana

#### Why People Use It?

- The expectation of a more intense high
- Easy access
- Belief it is safe
- Avoid detection



#### Tetrahydrocannabinol

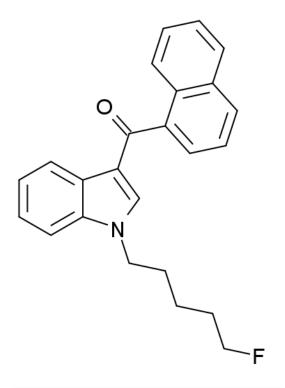
Binding affinity for CB1 K: 40.7 nM

#### JWH-018

Binding affinity CB1: 9 nM

#### AM-2201

Binding Affinity for CB1: 1.0 nM



All lipid soluble and highly volatized V PACC

## Synthetic Cannabinoids

- Psychoactive effects within minutes lasting up to 8 hours
  - Euphoria
  - Psychosis (9.4%-38%)
    - Visual
    - Altered perception of time
  - Anxiety
    - Panic
  - Agitation (23.4%)
  - Memory changes
  - Sedation (13.6%)
  - Confusion (12%)



## Medical Consequences

- Cardiovascular
  - Chest pain
  - Tachyarrhythmia
  - Cardiotoxicity
  - Tachycardia (40%)
  - Hypertension (8.1%)
  - Dizziness (7.3%)
- Gastrointestinal
  - Nausea
  - Vomiting
  - Hyposalivation
  - Xerostomia
- Renal
  - Acute kidney injury
- Other
  - Seizures-status epilepticus
  - Somnolence, dilated pupils, brisk reflexes
  - Appetite changes
  - Stroke

## Tolerance, Withdrawal and Dependence

 Cravings, restlessness, NMs, sweating headache, nausea, and tremor



## Diagnosis and Treatment

- Gas chromatography-mass spectrometry at selected labs
- Treatment is supportive
  - IVF
  - Calm setting
  - Benzodiazepines
  - Antipsychotic drugs prn



#### Sinus Aromatherapy for the Bath.

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## SYNTHETIC CATHINONES (BATH

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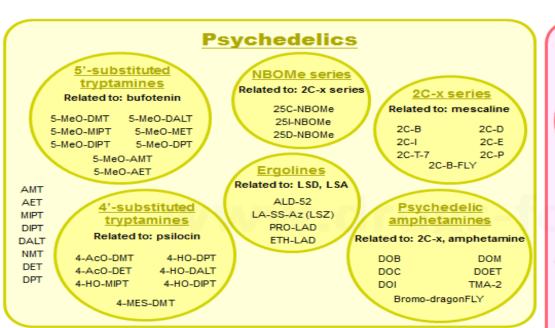
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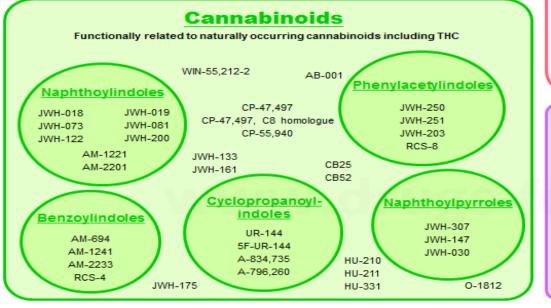
#### Stress Relief Aromatherapy for the Bath

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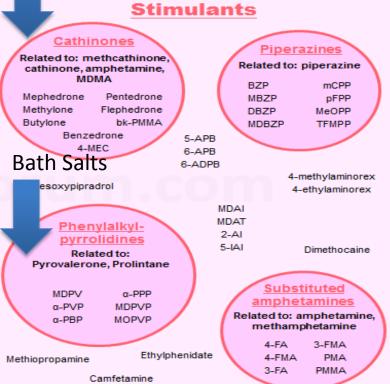
#### FILENCING NAMES

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#### Bath Salts



#### **Dissociatives**

Related to: ketamine, PCP

Methoxetamine
3-MeO-PCP
4-MeO-PCP
3-MeO-PCE
2-MeO-ketamine

#### Sedatives

Opioids

Related to: morphine, fentanyl, heroin α-methylfentanyl 3-methylfentanyl MPPP O-desmethyltramadol 7-acetoxymitragynine Metonitazene AH-7921

Phenazepam

## **COMMON BATH SALT NAMES**

- Meow meow
- Bubbles
- Cat
- Jeff
- Bliss
- Blue silk
- Cloud nine
- Drone
- Ivory wave

- Lunar wave
- Ocean burst
- Pure ivory
- Purple waves
- Red dove
- Snow leapard
- Stardust
- Vanilla sky
- White night



## BATH SALTS = SYNTHETIC CATHINONES

- Similar to other stimulants
  - Increase synaptic
     concentration of amines:

NE, DA, 5HT

amphetamine

cathinone

methamphetamine

mephedrone

Khat



#### Why People Use It?

- Easy access Internet head shops
- Avoid detection
- Potential for more intense high



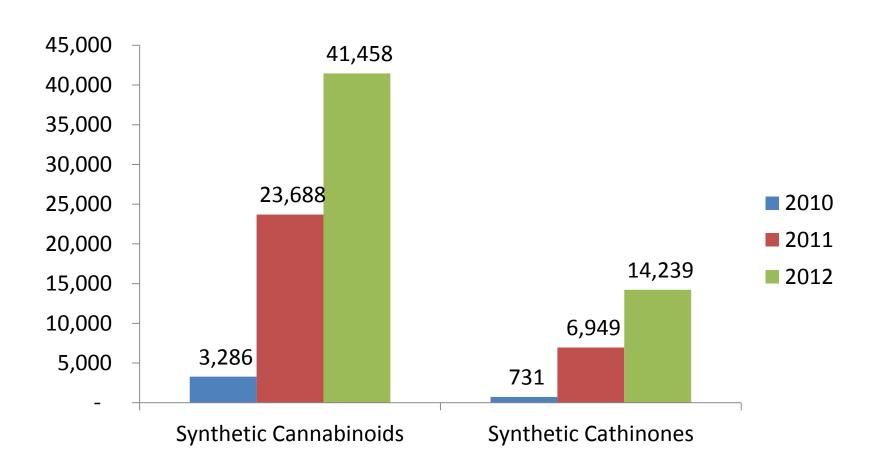
#### BATH SALTS = SYNTHETIC CATHINONES

- Typically white powder in small packets of 500mg-1gm for about \$25
- Average dose: few milligrams to >1gm
- Duration 2-3 hours
- Frequency 2-10+ per day
- Use
  - Nasal snorting
  - "Bombing"
  - "Keying"
  - Other: rectal, IV, IM, gingival, inhale





## EMERGING DRUG ITEMS IDENTIFIED IN U.S. NFLIS FORENSIC LABS: 2010-2012





# CLINICAL SYMPTOMS OF SYNTHETIC CATHINONE USE IN PATIENTS ADMITTED TO THE EMERGENCY DEPARTMENT

(N=236)

Agitation	82%
Combative/Violent behavior	57%
Tachycardia	56%
Hallucinations	40%
Paranoia	36%
Confusion	34%
Myoclonus/Movement disorders	19%
Hypertension	17%
Chest pain	17%
CPK elevations	9%



## CATHINONE: DIAGNOSIS

Clinical presentation

- Urine drug screen is not reliable
  - Even with specific screening most labs test only for Methylenedioxypyrovalerone (MDPV)
  - Sometimes false positive for methamphetamine
- History from patient and collateral sources



## CATHINO ONE: TREATMENT

- Supportive
  - Cardiac, Renal, Hyponatremia may necessitate hospitalization
  - Psychiatric may necessitate hospitalization for safety
    - Agitation: Benzodiazepines
    - Hallucinations: low-dose antipsychotics for hallucinations (Risperidone 1mg)
      - Avoid aggressive use of antipsychotics due to increased morbidity
- Monitor for vitals/lab abnormalities
  - Hyperthermia, dehydration, Renal function





## MDMA (ECSTASY)

- 3, 4-methylenedioxy-methamphetamine
- Street terms: Adam, E, X, XTC, love drug, Molly
- Cost per pill ranged from \$3-\$30
- Synthetic psychoactive drug
  - Both stimulant and hallucinogenic properties similar to methamphetamine and mescaline
- INCREASES DA, 5-HT, and NE
  - Inhibits 5-HT, DA, and NE reuptake, reverses transporters, decreases MAO activity
  - Releases 1.5 times more 5-HT than DA, more potent as 5-HT transporter inhibitor
- Releases oxytocin via directly and indirectly agonizing 5-HT→entactogenic



## WHAT IS "MOLLY"

- Ecstasy pills with little MDMA and lots of caffeine, meth, assorted drugs? <u>OR</u>
- A pure crystalline form of MDMA, most often sold as a powder filled capsule? OR
- 3. Methylone/Bath salts?
- Reports of desired effects of euphoria, but also increased paranoia, agitated delirium, scary hallucinations, psychotic episodes, violent or destructive self-harm behavior, including death
- Molly usually is not a pure form of MDMA, but may be a drug that can be very dangerous since its contents are unknown

## **MDMA (ECSTASY)**

- Adverse effects:
  - Serotonin Syndrome
  - Sweating, hyponatremia
  - Decreased coordination
  - Jaw clenching
  - Agitation
  - Hyperthermia
  - Rhabodmyolysis
  - hepatotoxicity

#### Mortality

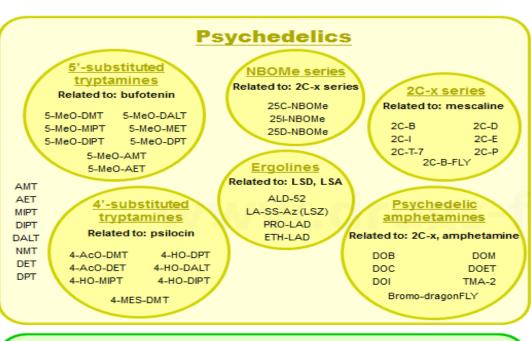
- 1997-2007 in the UK there were 605 deaths related to MDMA
- Polydrug use a large factor
- Hyperthermia related: 40-43°C (104-109), multiple organ failure
  - Induces apoptosis in cultured liver cells, acute liver failure
- Cardiac arrest
- Seizure
- Rhabdomyolysis

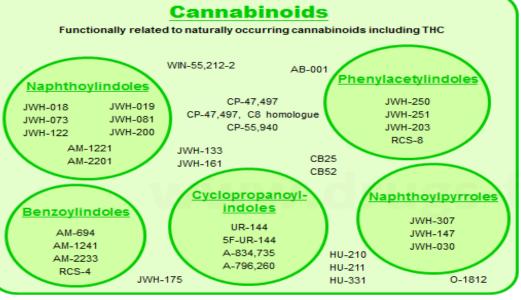


## MDMA Diagnosis & Treatment

- Serum and urine assays for MDMA
  - Negative test cannot confirm symptoms are result of MDMA toxicity
- Treatment for toxicity
  - FD Referral
  - Symptomatic care
    - IVFs, Oxygen
    - HTN-benzos 1<sup>st</sup> line and usually sufficient
    - Agitation-benzos
    - Psychosis-
  - Monitor
    - Serial neurological checks
    - Telemetry
    - Labs-Glucose, Renal/Liver
    - Urine output
  - Cool core temperature
  - Seizure treatment with benzodiazepines
  - Activated charcoal for ingestion within 1 hour







#### **Stimulants**

#### Cathinones

Related to: methcathinone, cathinone, amphetamine, MDMA

Mephedrone Pentedrone Methylone Flephedrone Butylone bk-PMMA

Desoxypipradrol

Benzedrone 4-MEC

...20

#### **Piperazines**

Related to: piperazine

 BZP
 mCPP

 MBZP
 pFPP

 DBZP
 MeOPP

 MDBZP
 TFMPP

5-APB 6-APB 6-ADPB

4-methylaminorex 4-ethylaminorex

Phenylalkylpyrrolidines

Related to: Pyrovalerone, Prolintane

> MDPV α-PPP α-PVP MDPVP α-PBP MOPVP

Methiopropamine Ethylphenidate

Camfetamine

MDAT 2-AI 5-IAI

MDAI

Dimethocaine

Substituted amphetamines

Related to: amphetamine, methamphetamine

> 4-FA 3-FMA 4-FMA PMA 3-FA PMMA

#### **Dissociatives**

Related to: ketamine, PCP

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3-MeO-PCE 2-MeO-ketamine

#### **Sedatives**

#### Opioids

Related to: morphine, fentanyl, heroin

α-methylfentanyl 3-methylfentanyl MPPP

O-desmethyltramadol 7-acetoxymitragynine

> Metonitazene AH-7921

Phenazepam

## **PIPERAZINES**

#### Street Names:

- "Frenzy"
- "Charge"
- "Herbal ecstasy" or "Legal X"
- "A2"

#### Two classes:

benzylpiperazines (BZP) and phenylpiperazines (TFMPP)

- Mimics effects of ecstasy (MDMA): 15-HT, DA, NE
  - Particularly dangerous if mixed: seizures and coronary syndromes

#### **Adverse Events**

- HTN/Tachycardia
- Reduced consciousness
- Psychosis
- Hyperthermia/coma



## **DEXTROMETHORPHAN (DXM)**

- Dissociative hallucinations
  - distance from reality
  - visual effects
  - perceptual changes
  - mystical-type experiences similar to psilocybin.
- Tachycardia, hypertension, agitation, ataxia, and psychosis at high doses; need to be careful of serotonin syndome
- OTC often found in combination
  - Acetaminophen
  - Diphenhydramine
  - Phenylephrine or pseudoephedrine
  - Alcohol
- Monitoring the Future survey 2011
  - 5% of 12 graders used in the past 12 months



## **DEXTROMETHORPHAN (DXM)**

- Street names include
  - "Robo" or act of using "Robo-tripping"
  - Triple Cs
- "Plateaus"
  - Darkridge.com street lore regarding different levels of high
    - "1st plateau (100-250mg): Great for conversation, yet can be a powerful trip. It all depends on the mindset. Usually one is very empathetic, and has great insights to everything"
    - "2nd plateau (250-450mg): All the effects of the 1st plateau are present, with the addition of more euphoria, a decreased sense of time and a decreased sense of surroundings."
    - "3rd plateau (450-800mg): Many of the effects of the first two plateaus are present + Visual (closed-eye-visual, mostly) hallucinations. Insights are extraordinary. "
    - "4th plateau (800-1800mg): The highway can be crossed at this plateau. What is on the other side of the highway could be anything. Alien encounters, out of body experiences, etc. are not uncommon."



## "SYRUP"OR SIZZURP

- Codeine + Promethazine cough syrup
- Cut with jolly ranchers/flavoring, and soft drink
- Sedation, pain relief, euphoria
  - Promethazine potentiates the codeine
- Hip-Hop/Rap music phenomenom
  - Pimp C death 2008 associated
  - Lil Wayne continues to struggle

## GAMMA HYDROXYBUTYRATE (GHB)

- 2 types of users
  - Body builders, self-med: depression, anxiety, insomnia
  - Recreationally: clubs, socially enhancement of sexual experience/aphrodisiac
  - Will use concurrently with MDMA, cocaine, alcohol, methamphetamine
  - Higher use among young club goers



## GAMMA HYDROXYBUTYRATE (GHB)

#### Acute Intoxication

- Unpredictable product, steep dose-response curve → difficult to titrate dose and overdose is common
- Vitals: hypotension, bradycardia, decreased respiratory rate, hypothermia
- CNS: Coma (within 15 minutes)
  - Agitation: including self-injurious behaviors
  - Amnesia
  - Seizure/seizure-like: myoclonus,
  - Ataxia, disconjugate gaze
- Respiratory depression
- CV: brady and hypotension, arrhythmias
- GI: vomiting, incontinence





# WHAT DO YOU DO IF SOMEONE HAS TAKEN AN UNKNOWN DRUG?

- Call your local poison center at 1-800-222-1222
  - 57 poison centers around the country have experts waiting to answer your call.
  - The experts at the Center can help you decide whether someone can be treated at home, or whether he or she must go to a hospital.
- Dial 9-1-1 immediately if they:
  - Stop breathing
  - Collapse
  - Have a seizure

...or if they have taken one of these and are having physical symptoms or behaving in a way that is concerning to you

#### For More Information

http://www.teens.drugabuse.gov/

http://www.drugabuse.gov/drugs-abuse/club-drugs

http://drugpubs.drugabuse.gov/promotions/back-to-school

http://www.samhsa.gov/



# SPECIAL THANKS – MARK DUNCAN, MD



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