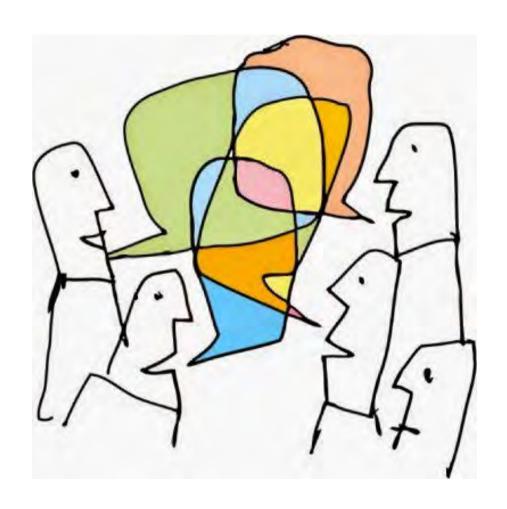
Substance use and HIV Scope of the problem

Ignacio Pérez Valero Hospital Universitario La Paz (Madrid)

What is the definition of substance use?



• Substance use:

Ingestion of psychoactive substance in moderate amounts that does not significantly interfere with functioning

Main categories of psychoactive substances

Depressants:

Result in behavioral sedation (alcohol, anxiolytics or hypnotic drugs)

Stimulants:

• Increase alertness and elevate mood (cocaine, amphetamines, nicotine)

Opiates:

Primarily produces analgesia an euphoria (heroin, morphine, codeine)

Hallucinogens:

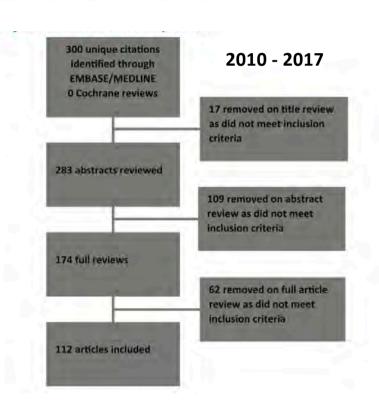
Alters sensory perception (marihuana, LSD)

Is it possible to scope the problem of substance use and HIV or are we in front of an impossible mission?



Sexualised drug taking among men who have sex with men: a systematic review

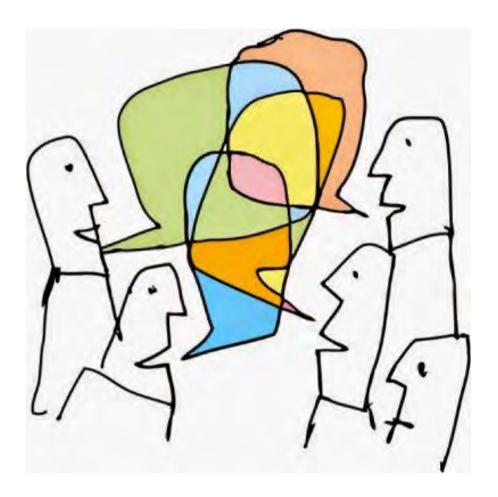
Just a brief example to understand the difficulty of the task



Recreational drug	Number of citations reporting usage	Range of reported prevalence of drug use in general MSM community	Range of reported prevalence of drug use in MSM in sexual health setting 1.2%–100% 2%–88.7%	
Methamphetamine	4411.12.14-54	4.0%-100%		
Poppers/nitrates	425,10,12,14-17,20,24,26,30,32,34-40,44,46,49-52,55-71	3%-37.3%		
GHB/GBL	411,2,5,11,13,17,18,20,26,28,32,34,38,39,43-46,49,51,58,62,63, 65-67,69,70,72-84	<1%-100%	1.4%-67.4%	
Crystal methamphetamine	81.2.5,10,13.24,30,44,56,58,60,61,63,65,66,68-70,75-92 <1%-100%		0.6%-100%	
Cocaine	342.5,11,12,15,27,32,34-39,41,43,49,51,59,61-66,70,72,74,75,85,87,90,93	5.0%-63.6%	1%-100%	
Erectile dysfunction medications	185,10,12,18,34,36,37,39,44,49,55,57,59,64,66,77,85,93	3.0%-33.1%	0.3%-56%	
Ketamine	252,5,14,17,24,26,34,39,40,44,46,49,51,55,57,61,62,65-67,70,72,75,77,84	0.8%-5.2%	1.3%-48.3%	
Ecstasy	3310-12,24,30-34,37-40,46,51,52,57,61-70,74,75,77,83,93,94 1,0%-38.3%		2.0%-68.3%	
Amphetamine	1413,24,27,33,40,46,49,55,59,63,66,70,77,95	0.7%-25%	0.2%-11%	
Mephedrone	261.2.5.15.28.39,45.46.49.58.62.66.67,70,72,75-84,93	1.3%-100%	3.0%-83.7%	
Cannabis/marijuana	1915.31.34.36.39.42.44,52,56,60-63,66,69.70,77,79,96	7%-54.8%	1.0%-56%	
Other drugs	26ª	0.6%-40.5%	0.2%-82.4%	

Tomkins A et al. Perspect Public Health. 2018:1757913918778872.

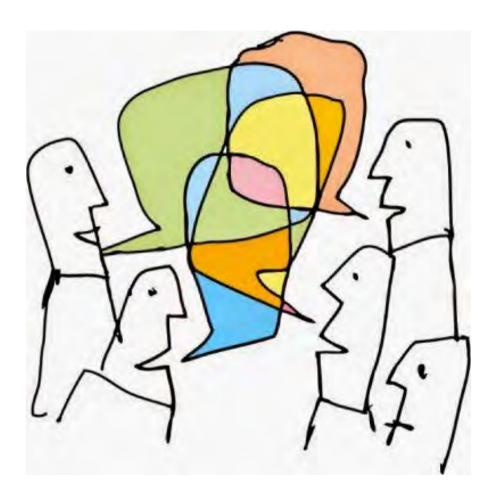
Why are these wide differences?



Substance use:

Ingestion of psychoactive substance in moderate amounts that does not significantly interfere with functioning

- Do you all have the same concept of substance?
 - Cocaine, heroin, crystal, mephedrone

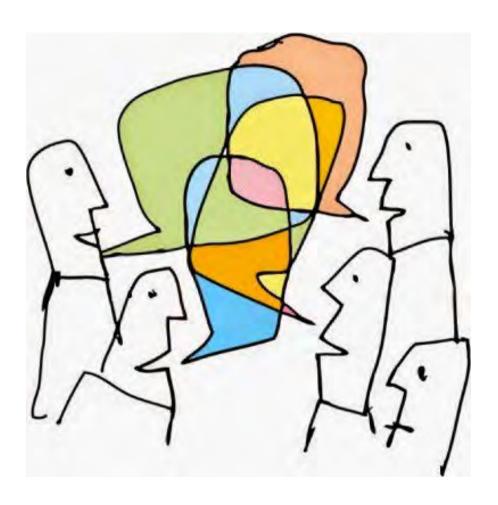


Substance use:

Ingestion of psychoactive substance in moderate amounts that does not significantly interfere with functioning

• Do you all have the same concept of substance?

- Cocaine, heroin, crystal, mephedrone
- What about poppers or viagra?

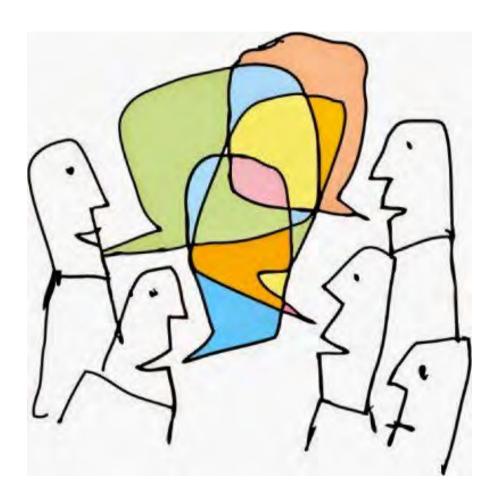


Substance use:

Ingestion of psychoactive substance in moderate amounts that does not significantly interfere with functioning

• Do you all have the same concept of substance?

- Cocaine, heroin, crystal, mephedrone
- Poppers or viagra
- What about tobacco or alcohol?



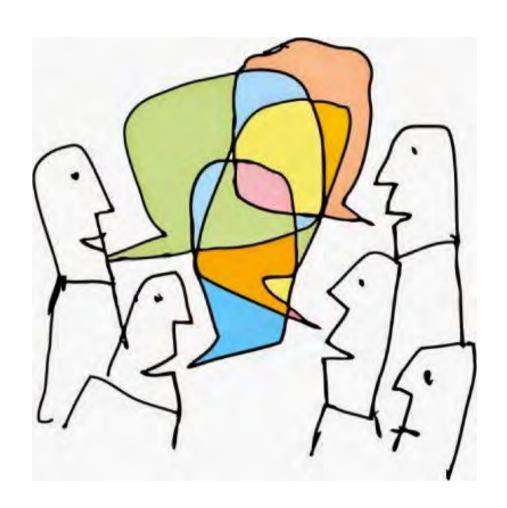
Substance use:

Ingestion of psychoactive substance in moderate amounts that does not significantly interfere with functioning

• Do you all have the same concept of substance?

- Cocaine, heroin, crystal, mephedrone
- Poppers or viagra
- Tobacco or alcohol
- And what about coffee?

Second limitation: Substance use, abuse and dependence



Different concepts that overlap in the literature

Substance use vs. substance abuse

Substance use:

Intentional ingestion of psychoactive substance in moderate amounts that does not significantly interfere with functioning

Substance abuse:

Compulsive (uncontrolled) use of one or more psychoactive substance that interfere with functioning **Definition**: Maladaptive pattern of substance use leading to clinically significant impairment or distress, as manifested by:

- Recurrent substance use resulting in failure to fulfill role obligations at work, school or home
- Recurrent use in physically hazardous situations
- Recurrent substance-related legal problems
- Continued use despite social or interpersonal problems caused by the substance
- Does not meet criteria for substance dependence

Steps leading from substance use to substance abuse

Abuse

Using drugs in an uncontrolled and excessive way

Regular use

High risk of progression Using drugs on a regular basis but moderately

Occasional use

Using drugs once in a while (less than once a week)

Experimental use

Trying one or more drugs

Substance addiction

Substance addiction

Chronic medical disorder with or without physiological dependence that includes multifactorial genetic components, biologic changes due to exposure to addictive substances and behavioral components.

<u>Definition</u>: Three of more of the following in the same 12 month period:

- Tolerance: Need for use of increasing amounts of the substance in order to achieve intoxication
- Withdrawal symptoms typical for the substance
- Substance taken in larger amounts or over a larger period of time than intended
- Desire to cut down or control use
- Great deal of time spent on using, obtaining, or recovering from the substance
- Reduced social, occupational or recreational activities because of substance use
- Continued use despite adverse physical or psychological consequences

Substance abuse vs. addiction



SUBSTANCE ABUSE CAN APPEAR CASUAL, WHILE ADDICTION USUALLY EXHIBITS MORE SERIOUS SYMPTOMS





#2

ADDICTION IS NEARLY ALWAYS ASSOCIATED WITH WITHDRAWAL SYMPTOMS



ADDICTION IS CONSIDERED A MENTAL DISORDER OR DISEASE, WHILE SUBSTANCE ABUSE IS NOT NECESSARILY A DIAGNOSIS

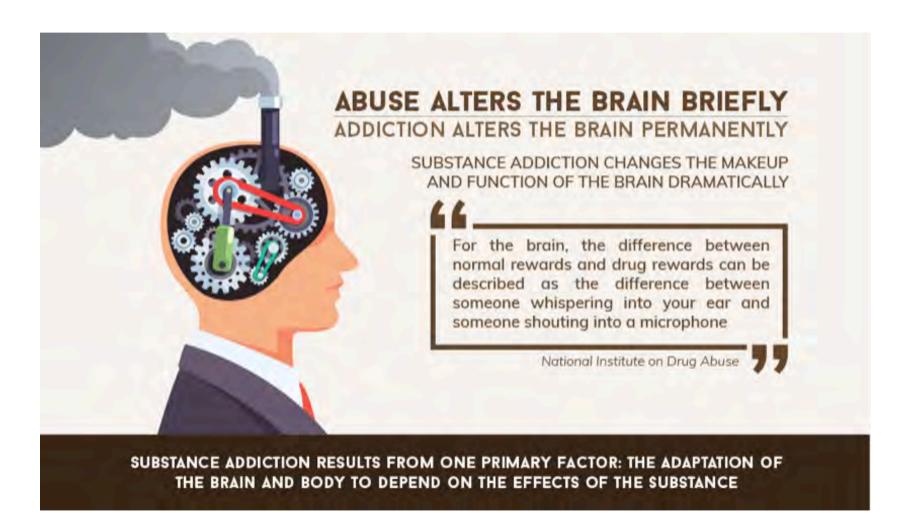






SUBSTANCE ABUSE ALTERS THE BRAIN BRIEFLY, WHILE SUBSTANCE ADDICTION ALTERS THE BRAIN PERMANENTLY

Substance abuse vs. addiction



What is the connection between substance use and HIV?

Causes of substance use are common in HIV+ individuals

- Environment of drug use
- Inability to resist peer pressure
- Availability of drugs
- Desire to look "cool", macho or grown-up
- Positive attitudes toward drug use
- Curiosity
- Belief that drugs will help overcome feeling of sadness, loneliness, boredom, anxiety
- Advertisements, media, apps

Substance use increase the risk of HIV acquisition

- Substance abuse is a primary vector for spread of HIV
- Substance abusers are more likely to engage in high risk behaviours
- Patients with substance use disorders may not seek health care
- Addiction and high-risk sexual behaviours have been linked across a wide range of settings

Substance use increase the risk of HIV acquisition: Epidemiology

- Outside the sub-Saharan Africa, 10% of all new HIV infections are attributed to IDU despite significant regional variations
- According to the WHO-HIV/AIDS program, IDU is responsible of more than 80% of all HIV infections in eastern Europe an central Asia
- The epidemic in countries in the middle East and North Africa have been largely attributed to IDU, and it is currently linked to the growing epidemic in Indonesia, Vietnam and Malaysia.
- In the US drug abuse is a significant risk factor for HIV
 - 1 million people in the US are living with HIV and about 1/3 of these cases are linked directly or indirectly to IDU
 - Up to 25% of all new diagnoses in US are reported in IDU

Source: Drug abuse and HIV/AIDS, a research update from the NIDA

Some data of substance use from the US

The alcohol use problem among US general population

- More than 50% of adult US population drank alcohol in the past 30 days.
- 17.3 million Americans (5% of US population) are heavily drinkers.
 - 29.4% of them are also current illicit drug users
- Costs of alcoholism and illicit drug abuse to the US economy
 - Productivity, health and crime costs of alcoholism and drug addiction: \$235 billion
 - US business cost of alcoholism and drug addiction: \$120 billion
 - Excessive alcohol use is the 3rd leading life-style cause of death

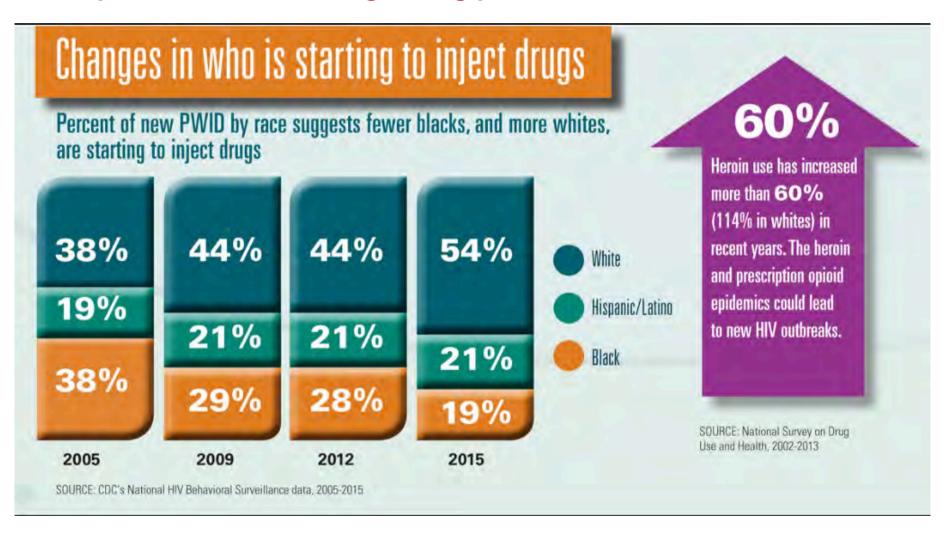
Source: National Institute of Drug Abuse

The substance use problem among US general population

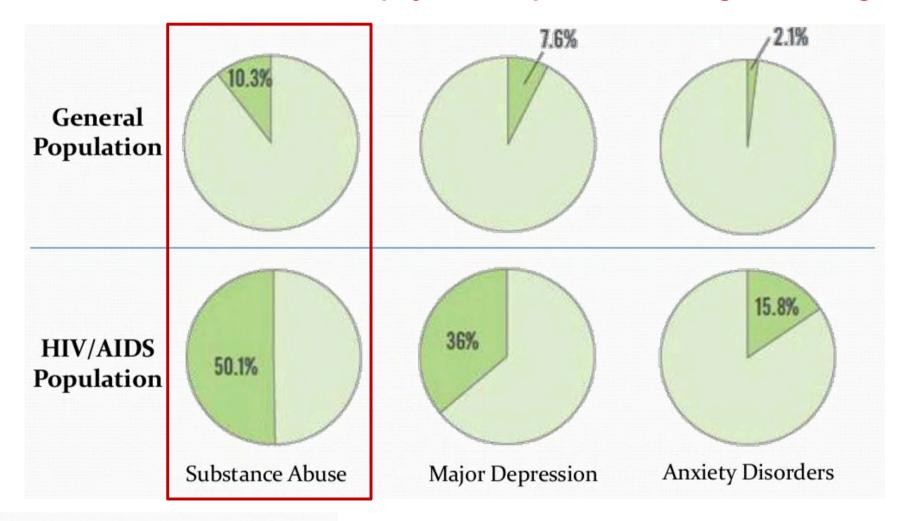
- 20.1 million Americans aged 12+ are current users of illicit drugs (8% of all the population)
 - This prevalence remains stable since 2002
- 15.2 million Americans aged 12+ consume cannabis
 - 57.3% of them only consume cannabis
- 8.6 million Americans aged 12+ consume other illicit drugs
- 6.2 million Americans aged 12+ consume non-prescribed psychotherapeutic drugs
 - 4.7 million consume non-prescribed pain-relievers
 - 1.8 million consume non-prescribed tranquilizers
 - 0.9 million consume non-prescribed stimulants
 - 0,2 million million consume non-prescribed sedatives

Source: National Institute of Drug Abuse

IV heroin epidemic in the US, a growing problem

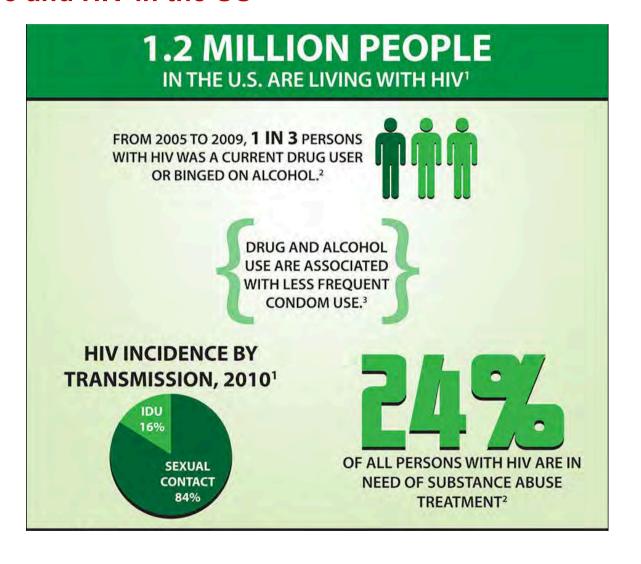


Prevalence of substance use and psychiatric problems is higher among HIV+



Source: BW Hospital, Harvard Medical School, 2012

Substance use and HIV in the US



National (US) HIV Behavioral Surveillance (2014-2016) Survey – non-MSM subset

Type of study: CDC Survey

Target population: Low income non-MSM from 17 US cities

Timeline: 2016

N = 7,560



Substance use:

Crystal meth use HIV-: 4% (injectable: <1%)

Crystal meth use HIV+: 4% (injectable: <1%)

Risk factors / behaviors (last year):

Any STI (last year): 6.9% (HIV+ 10.3%)

• Female: 8.2% (vs. male: 5.4%)

• HIV testing (last year): 40.5%

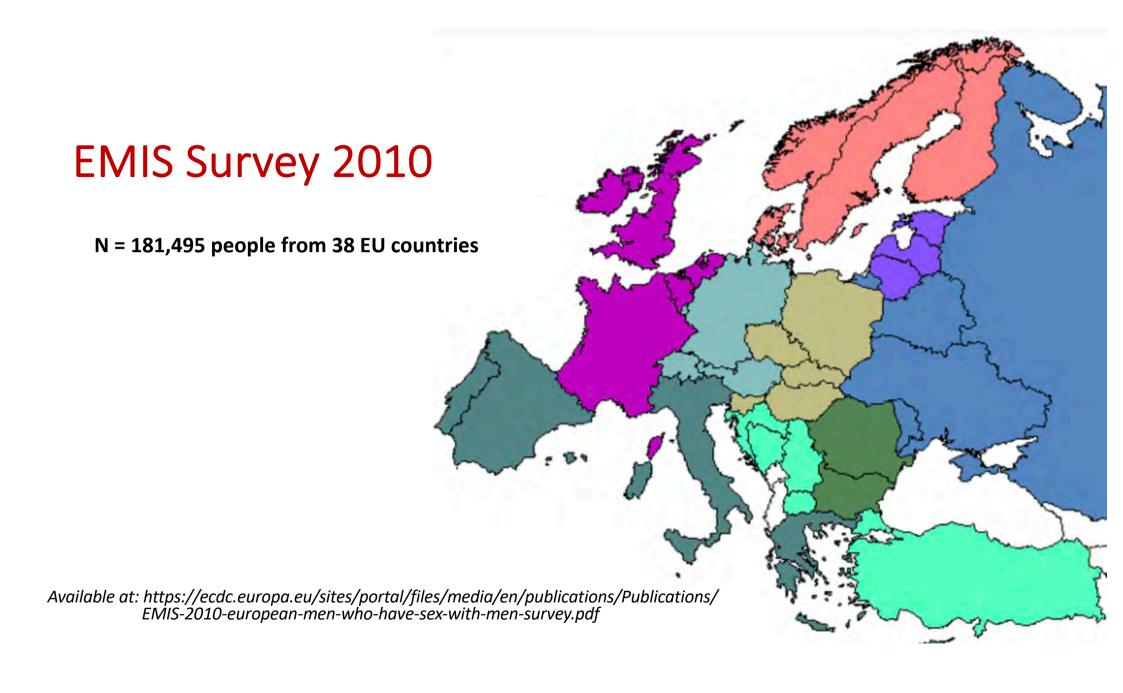
PrEP use: 0.2%

• Condomless sex (last year):

• Men: 45%

• Female: 40%

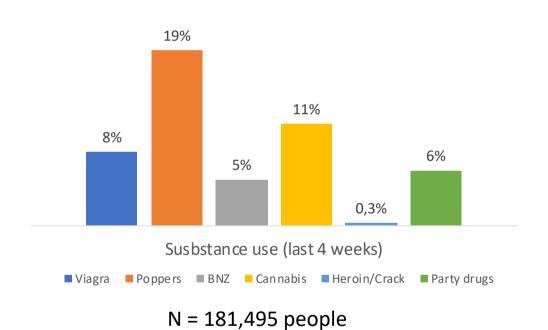
Some data of substance use from EU

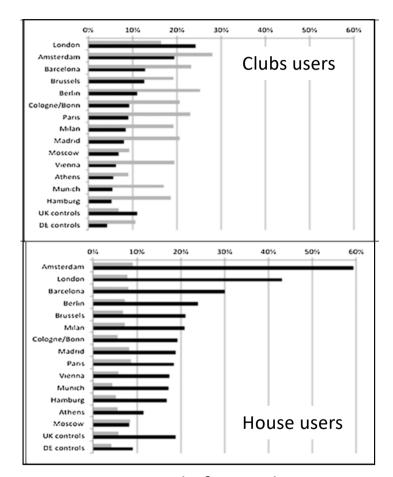


EMIS Survey 2010 – Demographics

- Men: >99% (<1% trans). Age <30 years: >50%
- Lived in medium/small cities (<500,000 people) or in the countryside: 54%
- Highly educated men: 60% (30% 83%)
- Unemployed: 6% (0.8% 11%)
- MSM: 76% (+15% bisexual)
- Steady relationship with a man: 39% (54% were single)
- Living alone: 30% (13% 50%)
- Apps use in last 7 weeks: 94%
- Sexual unhappiness among MSM: 46.7% (27.8% 61.3%)
 - Mean reason: "I want a steady relationship": 25.4%

EMIS Survey 2010 – Substance use





N = 55,446 people from urban centers

Available at: https://ecdc.europa.eu/sites/portal/files/media/en/publications/Publications/Publications/EMIS-2010-european-men-who-have-sex-with-men-survey.pdf
Schmidt AJ et al. Int J Drug Policy 2016;38:4-12.

EMIS 2010 Substudy: Spain

Type of study: EMIS survey subanalysis

Target population: MSM app users

Timeline: June – August 2010

N = 13,111



Chemsex use (last year):

• Chemsex use (last year): 21.9% (included other drugs)

• Sex enhancers (last year): 33.6%

Risk factors / behaviors (last year):

• UAI (last year): 29.4%

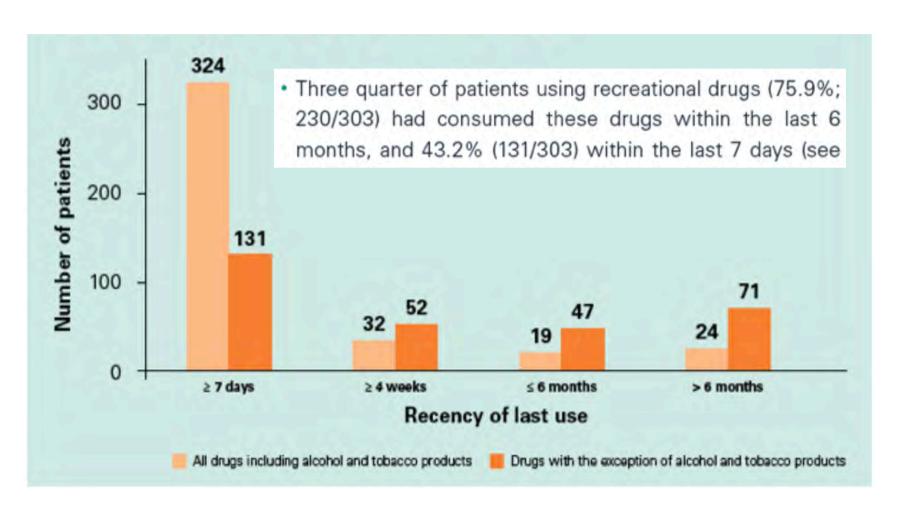
• HIV+ with VL > 50 cop/ml or unknown: 27.8%

• STIs: not reported

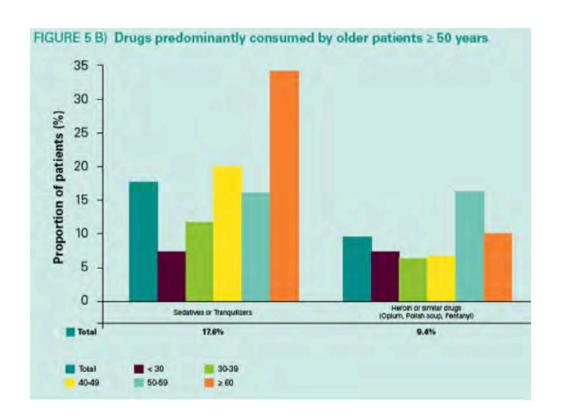
• HIV+: 12%

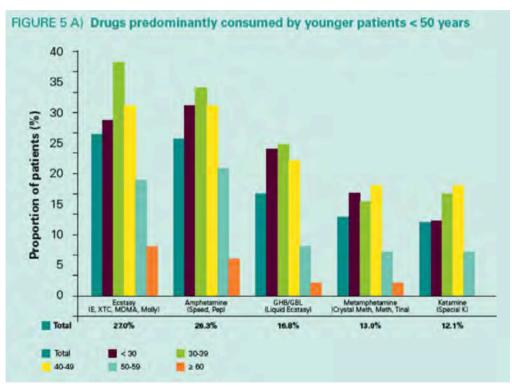
Never HIV tested: 26.2%

Proportion of drug users among German HIV+ individuals



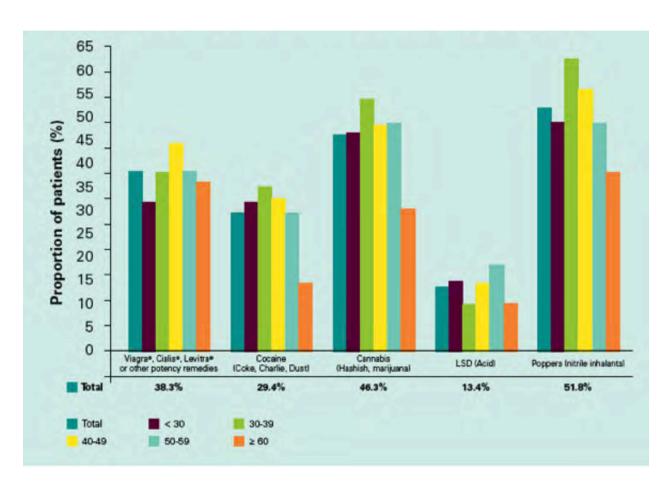
Drugs consumed by German HIV+ individuals by age





- Olders: sedative drugs and Morphine-like drugs
- Youngers: Ectasy, GHB, metamphetamine, ketamine, amphetamine.

Drugs consumed by German HIV+ individuals without preference by age



• Viagra, cocaine, cannabis, LSD, Poppers

USEX Study (Madrid, Spain) – MSM HIV+ Survey

Type of study: Self-reported survey

Target population: MSM HIV+ at 22 outpatient clinics

Timeline: June 2016 - March 2017

N = 742



Chemsex use (last year): 29.1%

• GHB/GBL: 21%

• Mephedrone: 20.1%

• Crystal Meth: 8.7%

• Slamsex: 4.6%

Risk factors / behaviors:

VL >50 (last month): 34% (vs. 17% non-drug users)

• UAI (last 6 months): 87% (vs. 43% non-drug users)

• Fisting (last 6 months): 44% (vs. 6% non-drug users)

>20 sexual partners (last 6 months): 44% (vs 8%)

Any ITS (ever): 85% (vs. 53% non-drug users)

• Depression: 33% (vs. 21% non-drug users)

Gonzalez-Baeza A. et al. AIDS Patient Care and STDs 2018;32:112-118

Some data from other countries

Hanoi (Vietnam) HIV/AIDS prevention center study

Type of study: Peer to peer interview

Target population: MSM (hidden population)

Timeline: Sept – Dec 2014

N = 622



Substance use (ever):

• Crystal Meth: 20.1%

• Other sexualized drugs (ever):

• Poppers: 9.7%

• Erectile dysfunction medication: 8.5%

• No Slamsex reported

Risk factors / behaviors:

• Involved in selling sex (AOR: 2.43)

Facilitate social MSM networking (AOR: 3.23)

• No ITS rates reported

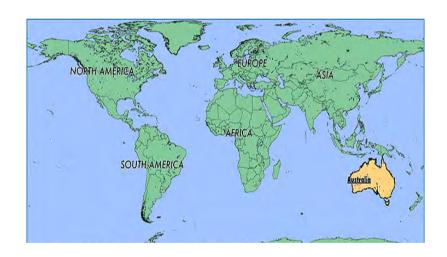
The FLUX Study (Australia)

Type of study: Online questionnaire

Target population: MSM apps users (Facebook 65%)

Timeline: Sept 2014 – July 2015

N = 2,251 (1711 had 6-month follow-up)



Substance use (last 6 months): 28%

• Crystal Meth: 12%

• GHB/GBL: 6.9%

• Any drug users (last 6-months): 50.5%

• Poppers: 32.1%

• Erectile dysfunction medication: 21.8%

• Slamsex (last 6 months): 4.7% (ever 10.3%)

Risk factors / behaviors:

>10 sexual partners (OR: 3.21)

• UAI (OR: 1.81)

• HIV+ (46.2% vs. 5%)

• HCV (16.1% vs 1.2%)

Consequences of substance use (in HIV)

Consequences of substance abuse in HIV+ individuals: Comorbidities

- The accumulation of medical sequelae from chronic substance abuse can amplify the burdens of the HIV infection itself and conduct to an accelerated aging process
- HIV+ substance abusers become vulnerable to infections (hepatitis, ITS, sepsis, soft tissue infections, endocarditis...)
- Neurological symptoms may overlap between HIV infection and substance abuse
- Drug users have a higher risk of developing depression than non-drug users
- Drug use was associated with 4.1 times greater risk of being a poor adherer (<90%), specially among stimulant drug users
- Drug users have an strong stigma
- Lower rates of linkage to care

Consequences of substance abuse in HIV+ individuals: DDI

TABLE 1: Proportion of patients consuming recreational drugs with a potential of drug-drug interactions some any antiretroviral therapies

Viagra®, Cialis®, Levitra® or other potency remedies	38.3%	Ecstasy (E, XTC, MDMA, Molly)	27.0%	Heroin or similar drugs (Opium, Polish soup, Fentanyl)	9.4%
Cocaine (Coke, Charlie, Dust)	29.4%	Amphetamine (Speed, Pep)	26.3%	Mephedrone or similar drugs (Bubbles, M-CAT, Meow, Meph)	5.8%
Sedatives or Tranquilizer	17.6%	GHB/GBL (Liquid Ecstasy)	16.9%	Alcohol	91.1%
Anabolic steorids	6.0%	LSD (Acid)	13.5%	Poppers (nitrile inhalants)	51.8%
Psychoactive substances (Legal Highs, Herbals)	5.1%	Metamphetamine (Crystal Meth, Meth, Tina)	13.0%	Tobacco products	74.5%
Cannabis (Hashish, marijuana)	46.3%	Ketamine (Special K)	12.1%		4-4-

Consequences of substance abuse in HIV+ individuals: ART deferral

