



#### **Background and Introduction**



Drug abuse of psychotropic agents is an important medical and social problem which affects 7000-10000\* individuals in HK

Drug abusers who develop seizure disorders constitute a serious health problem with many unmet needs

The Division of Neurology, Department of Medicine and Therapeutics, Prince of Wales Hospital has long recognized the importance of psychotropic drug use and seizure disorders/ epilepsy.











A study in 2011 already identified ketamine, methamphetamine and cocaine as important substances of abuse in this region\*. Other drugs of abuse are also included in the current study.

Among psychotropic drug abusers, apart from psychiatric comorbidities, seizures are often overlooked

The mechanism of seizures can be toxic overdose, or following drug withdrawal.

The latter is particularly important as drug abusers may run out of medications due to financial or other constraints.

Our study will explore whether prescriptional drugs are subject to abuse.

We want to understand whether treatment of seizure should be hand-in-hand with detoxification. Could there be long-term sequelae of seizures even after detoxification?





## The clinical question and need %

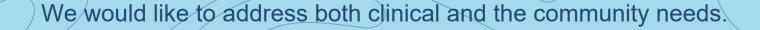


General doctors usually find it difficult to serve these clients. Why? These patients get admitted to the medical wards rather than psychiatric wards as *obiter* patients but we often lack the special team to help with these patients



We would like to thank BDF for the promulgation of this project and greatly helping the clients in need.

There is a great opportunity for collaboration and training with this project.





# The BDF project for the psychotropic drug abusers



HA: Dr H Leung (Chief of Neurology, Service) CUHK: Prof V Mok (Chief of Neurology, Academic Affairs) Part-time doctor: Dr CK Wong Full-time nurse: Ms Celia Tse







Clinical screening from medical wards/ neurology wards





Treatment of seizures and detoxification



Outpatient encounters from neurology clinics







### **Output indicators**



(1) Provision of 720 in-patient visits/ outpatient sessions→ 875 (120%)



(2) Serving 120 patients with seizures and psychotropic drug abuse → 114 (95%)



(3) Reaching 120 family members of psychotropic drug abusers

→ 111 (93%)



(4) Providing 3 outreach seminars for 100-200 secondary school students → 2 seminars and 1 webinar (100%) and 508 students (254%)

#### **Outcome Indicators\***



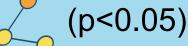


(76 patients had 12-month outcome and 108 patients had a variable period of 1-12-month outcome. 9 patients either cannot be contacted or have already passed away):

69.4% with improvement in seizures

62% achieving seizure freedom





\*subject to finalization and amendment before submission of end-of-project report



#### **Outcome Indicators**



#### (2) Rehabilitation outcome



- 78.7% of patients achieving antiepileptic agent compliance and
- 52.8% showed 0-1 symptoms in tolerability questionnaire
- 72% reported statistically significant improvement in quality of life (QOLIE-31) (p<0.001)\*

  (Wilcoxon signed rank test, n=77, sum of positive ranks 782, negative ranks 1993, z value -3.26, p<0.001)
- 84% reported reduction of drug use
- 67.5% reported quitting of drug use







#### **Outcome Indicators**





(3) Anti-drug attitude with medical consultations

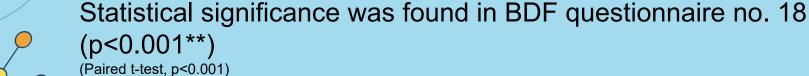
75.9% reported improvement in anti-drug attitude.

Statistical significance was found in BDF questionnaire no. 18

(p<0.001\*)

(Wilcoxon signed rank test, n=73, sum of positive ranks 344, negative ranks 2357, z value = -5.53, p<0.001)

(4) Anti-drug attitude with school seminars

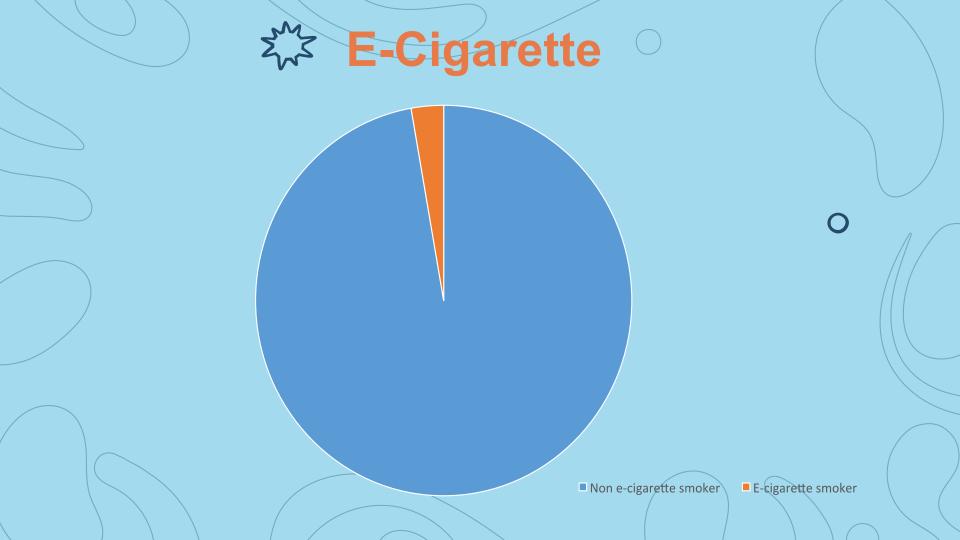




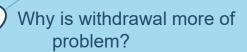
#### Types of psychotropic drug use



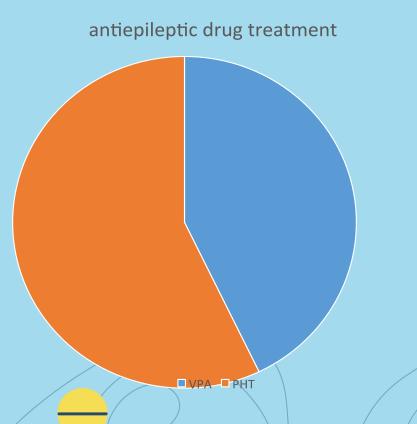


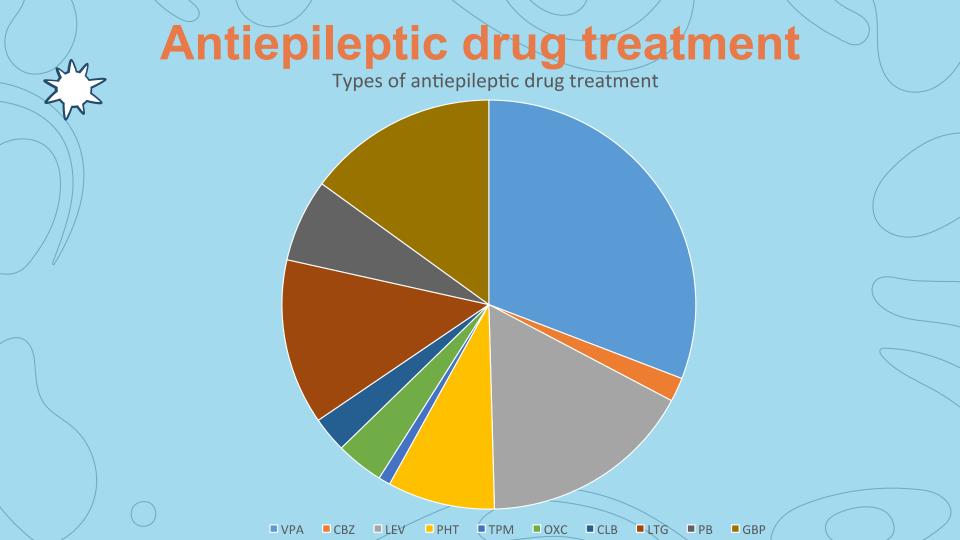


#### Proportion of overdose/withdrawal

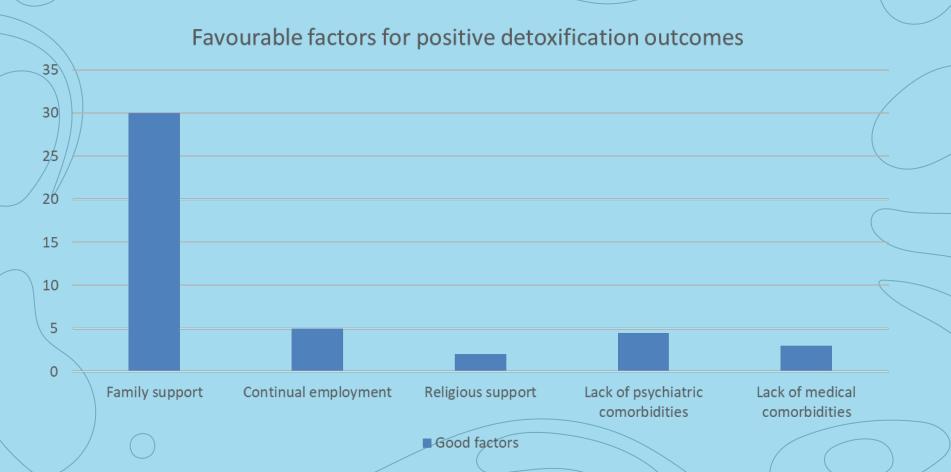


- 1. Frequency of events could be higher
- 2. Running out of medications is easier than taking an overdose
- 3. Overdose is more likely to enlist psychiatrist's support owing to possibility of suicidal attempt
- 4. Higher likelihood of seizure during withdrawal





#### Factors associated with detoxification outcome



# School talk to Ho Man Tin Government Secondary School

200

Before COVID, school talks were held as physical talks or

seminars



Lecture from doctor Dates: 14/9/2018 and 20/3/2019



Quiz proved to be highly popular

#### School talk to International School Foundation

- A school talk was successfully performed via the on-line format on 20<sup>th</sup> Jan 2021.
- Lecture delivered by Dr H Leung on psychotropic drugs and seizure
- Case sharing by 5 medical students
- Talk delivered by Ms C Tse on brain health
- Question and answer session







### How does COVID-19 affect us?

Suspension of clinic from Jan 2020 to Apr 2020 and again from Jul 2020 to Aug 2020.

Stopping clinical work by specialist nurse from Jan 2020 to Apr 2020 and again from Jul 2020 to Sep 2020.

The numbers were generally caught up after resumption of clinical duties as allowed by HA and CUHK. We will work hard with follow-up questionnaires and reach-out to family

\*\*\*\*

School talks can only be delivered by online format.



#### Zopiclone abuse and seizures

44 yr old English tutor

History of depression

Found by husband to have bathed for an abnormal duration

Drowsiness and brought to A&E

GTCS 5 min given PHT

Transpired that she had bromazepam and zopiclone from private practitioner

Intake was not entirely supervised

(4-5 tabs but up to 10 tabs if not able to sleep)

There was a period of withdrawal

Initially agreed by patient (pt) to go to psychiatric ward for detox

PHT changed to lamotrigine (LTG), folic acid

But then DAMA while in psych ward

(initially reluctant to enlist support from husband)

#### Zopiclone abuse and seizures

Another breakthrough seizure 1 week later

Finally admitted to psych for full detox

Motivational interview

Enlisted husband's support

Controlled replacement with zopiclone

(needs pill count)

No seizure >1 yr

Continual LTG as not able to entirely come off zopiclone

Anxiety with insomnia

Later on decided to emigrate

Letter for pt to receive continual care in foreign country

#### Tramadol abuse and seizures

40 yr old
Hx of IVDA, hep B/C and hx of ICH and epilepsy on VPA
Repeated admissions in 2019 and 2020 for seizures, one being status
Initially thought to have remote symptomatic seizure
Due to frontal encephalomalacia
VPA was titrated
Tramadol use only alleged to be 50mg qid

But informed by toxicology team that
Urine toxicology sample 3/20 showed supra-maximal peak suggestive of supratherapeutic exposure

#### Tramadol abuse and seizures

Pt all along declined psychiatric input
Multiple interhospital visits also created polypharmacy and abuse of
tramadol

- 1. Simplified medication follow-up visits to reduce multiple iatrogenic sources
- 2. Tackle the phlebitis and varicose vein
- 3. Goal orientated tapering
- 4. Phone follow-up by epilepsy nurse

No seizure for 7 months now Uses tramadol 3 times / week

Epidemiologic Reviews

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The ongoing opioid-overdose epidemic remains unabated.

DOI: 10.1093/epirev/mxaa002 Advance Access publication: April 3, 2020

In 2017, opioid-related deaths accounted for 68% of the total drug overdose deaths in the United States (1). From 1999 to 2017, the annual number of overdose deaths related to opioids (including licit and illicit opioids) increased more than 600% (1). Moreover, the types of opioids involved in these deaths have evolved. From 1999 to 2010, commonly prescribed opioids were the leading cause of opioid-related deaths; these leveled off from 2011 to 2016 and were surpassed by synthetic opioids as the leading cause of opioidrelated deaths in 2016 (1). Despite the evolution of the opioid epidemic from prescription to synthetic opioids as the leading cause of death, commonly prescribed opioids continue to play a significant role in the epidemic, and interventions to improve opioid prescribing behaviors remain a priority.

#### n Opioid-Related

erdá, Katherine M. Keyes,

iversity Mailman School of Public du).

#### Modafinil use and seizures

28 year old gentleman Works in big auditing firm History from family - he had been working under high pressure at work with lack of sleep lately 1st GTCS at home, called ambulance Another GTCS in ambulance, GCS 12, admitted directly to ICU Given iv phenytoin Denied any illicit drug use initially. No fever Further questioning – he said he was using some kind of "bodybuilding products/ vitamins/ private med"→ family asked to bring back packing Urine toxicology from ICU – pending

#### Modafinil use and seizures

Switched to oral phenytoin 300mg nocte

LP → CSF WCC 4 TP- n, HSV/VZV PCR -ve

Further seizures 2 days later, possible NCSE – given LEV, tided over with sedation

CPK up to 49000

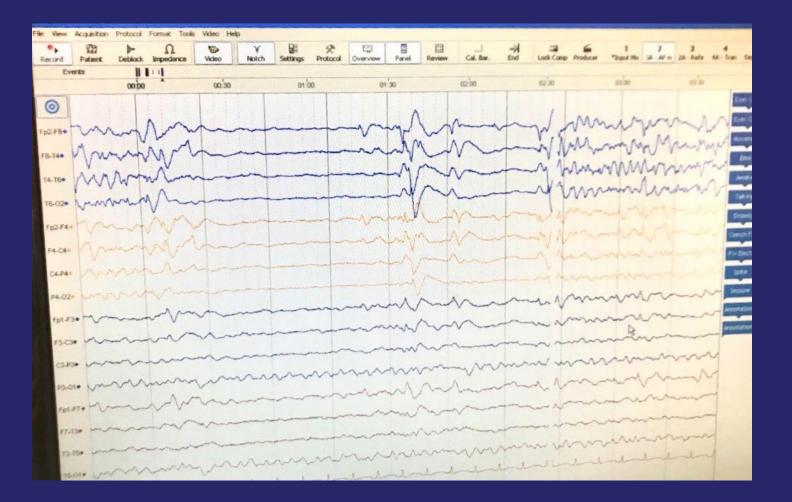
Family brought back some package which appeared to be medications from India and some vague words of "modafinil" and 30 tabs left behind EEG- alpha rhythm only

MRI brain – no obvious features of encephalitis

Later urine toxicology – only phenytoin

Pt had a few days of visual/auditory hallucination

Later admitted that every he and his colleague will work overnight one night / month with modafinil bought on-line



#### Modafinil use and seizures

Had liver derangement with PHT switched to GBP
Hep B/C neg and u/s abdo- n
Fleeting visual hallucination lasting a week
DDx – autoimmune encephalitis but pt subsequently improved
Oxford panel –ve
Urine toxicology -ve

6 months later – no seizure, LEV tapered off 12 months later – no seizure, GBP also tapered off



# Case sharing – refractory epilepsy and amitriptyline abuse

- 40 year old beautician
- Benzodiazepine dependence
- Amitriptyline abuse → cardiac arrests and other complications
- How did she get the amitriptyline through father in pharmacy
- Many admissions a year for seizure, mostly associated with amitriptyline abuse
- Not easy to find out if the seizure was due to withdrawal of benzodiazepine or overdose of amitriptyline



(con't)

# Case sharing – refractory epilepsy and amitriptyline abuse

- Multidisciplinary care with psychiatrists and toxicologists
- Due to the seizures, the patient was often admitted to medical ward
- On gabapentin, pregabalin, topiramate, valproate
- Replacement benzodiazepine from psychiatry
- Strategies

Short follow-up to ration medications Give her another "replacement" tablet Urine testing every time for amitriptyline

+ pill count

Piriton, from hospital



Amitriptyline, from pharmacy

## Case sharing – epilepsy developing long after the abuse has ceased

(8)

- 55 year old gentleman
- History of cough mixture abuse and intracranial haemorrhage ~10 years go
- Presented with GTCS and admitted to hospital
- Although there is no currently no more psychotropic drug abuse, a new onset epilepsy is diagnosed
- Started on phenytoin
- Remained seizure free but will likely require life-long medication.

#### Implications for drug abusers

- Seizures and epilepsy exemplified the scope of medical burden due to psychotropic drug abuse.
- It gives us an important message in our drug campaign
- Withdrawal from sleeping pills is by far the most important form of seizure due to psychotropic drug abuse
- Drug abusers lie somewhere in the borderzone area between medical doctors and psychiatrists
- May occur in all walks of life
- Psychiatric comorbidity, social problems and multiple medical problems are common
- Family support is an important indicator for success
- Epilepsy can occur even after the abuse has ceased

#### **Future directions**

- Our project has consolidated the role of nurses in this campaign
- A medical coordinator may yet fulfill the service gap
- Innovative approach between neurology, psychiatry is needed
- Success in dealing with psychotropic drug abusers is a win-win situation for HA
- More community work is needed
- COVID has not stopped people from having drug abuse
- But innovative ways to deliver care during times of COVID are necessary



