STUDY OF PATTERNS OF DRUGS OF ABUSE IN NEW TERRITORIES EAST AND WEST CLUSTERS DRUG OF ABUSE CLINIC USING CONVENTIONAL AND NEW TECHNOLOGIES

Final Report Submitted To

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By Research Team

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1 Introduction

1.1 Background

The rapidly changing pattern and increasing prevalence of abusing conventional and novel psychotropic substances in Hong Kong have resulted in a community-wide growth of concerns for better detection and control of the drug abuse problem. The purpose of developing accurate drug detection methods is to provide clinicians, therapists, and other parties that related to drug of abuse problems, an objective mean for diagnosis of illicit drug use and for treatment and monitoring of substance dependence.

Traditionally, a wide variety of drugs can be detected in urine specimens qualitatively by laboratory methods such as Immunoassay and High-Performance Liquid Chromatography with Ultraviolet Detector (HPLC-UV). However, these conventional methods of drug of abuse screening have many limitations including lack of sensitivity, specificity, and the requirement of a known drug library or pure standards for novel drugs detection that hinders the investigation, leading to mis-diagnosis and under-reporting of the problem.

A new technique was developed, namely, Ultra-Performance Liquid Chromatography Time-Of-Flight Mass Spectrometry (UPLC-TOF/MS), which allowed target screening and confirmation of conventional as well as novel drugs based on a combination of retention time, exact mass and fragmentation patterns of a drug molecule. This exact mass approach has the advantages of creating theoretical drug libraries from the molecular formula without the need of pure standard and of improving the sensitivity and specificity of compound identification that will greatly enhance the detection rate of conventional and novel drugs.

The Department of Chemical Pathology of the Chinese University of Hong Kong (CUHK), Prince of Wales Hospital (PWH) has newly installed such advanced equipment. With the collaboration of the Department of Chemical Pathology and the Department of Psychiatry of CUHK, PWH, a

study was conducted in the Drug of Abuse Clinic in PWH to investigate the pattern of drugs of abuse in the New Territories East (NTE) region of Hong Kong and to compare the difference between the conventional and the new techniques for the detection of illicit drugs in urine specimens.

1.2 Objectives

The primary objectives of our study are to validate the use of UPLC-TOF-MS by demonstrating the accuracy and reproducibility of drug detection through comparison of use of the conventional and new laboratory techniques and to develop prevalence data on current pattern of psychotropic substance abuse in the locality. In addition, comparing the drug abuse history obtained from subjects with respect to their corresponding urine analytical result to look for possible impurities and contaminants.

With the diagnostic utility of UPLC-TOF-MS, its use in clinical practice may help to guide the treatment of substance abuse. Effective preventive measures and specific treatment program targeting the problem may also be developed from the local data on preference of substances of abuse.

1.3 Study Design

In our study, we recruited patients attending the Drugs of Abuse Clinics of the PWH anonymously. Patients who were not willing to answer the questionnaire and/or to provide their urine samples were excluded from our study.

Sets of pre-numbered questionnaire (see section 7.1) and temperature-sensitive urine bottles were prepared for the anonymous recruitment of study subjects. The questionnaire focused on the demographics, socio-behavioural characteristics and the information of previous and last drug use of the recruited subjects. Twenty mL spot urine sample was collected from each subject and was aliquoted for measuring urine creatinine concentration and a panel of drugs of abuse using conventional immunoassay, HPLC-UV, and the latest UPLC-TOF/MS techniques.

1.4 Data collection

Due to the potential legal issue arisen if a positive urine result is obtained from subject who is under probation by law, the study subjects were recruited anonymously. All data collected from the recruited subjects was identified by a unique number generated by the laboratory and was kept confidential.

1.5 Difficulties encountered

Subject Recruitment

Response rate of subject recruitment was unexpectedly low ranging from 0 to 5 subjects per clinic session and thus the total recruitment was 52 subjects during the first year of our study. Despite the suboptimal response rate, the number of potential new cases recruited from the Drug of Abuse Clinic in PWH was also limited. Possibility of recruiting subjects from psychiatric in-patient wards or emergency department settings within the NTE cluster hospitals had been explored but was withdrawn due to the logistic difficulties.

To expand the source of recruitment, we collaborated with another project granted by the Beat Drugs Fund Association, "Socioeconomic and Health Impacts of Substance Abuse in Hong Kong – A Longitudinal Study", leaded by Professor Kenneth Lee from School of Pharmacy, CUHK. One of the objectives of this project is to study the acute toxicity in a group of drug abusers admitted or referred to a public hospital by performing a quality and reliable analysis on the urine specimens for toxicology screening. The Department of Chemical Pathology, CUHK, as one of the collaborating institutions, primarily provides the analytical services with both conventional technologies (HPLC & Immunoassay) and new technology (UPLC-TOF/MS). Relevant data obtained from the second project including the urine analysis result and information from questionnaire can be used by our first project for data analysis. The revised study protocol was approved and subsequent subject recruitment started since June 2008. With the expanded sources of subject recruitment from new cases of the Castle Peak Hospital (CPH) Substance Abuse Clinic (SAC) and various non-government organizations

(NGOs), the number of participants increased significantly.

Other measures to increase the recruitment include adoption of paying honorarium for subject recruitment and to recruit also the old cases of the SACs in the New Territories East (NTEC) and West (NTWC) Clusters Hospitals. The revised study protocol was approved by the Joint CUHK-NTEC Cluster Clinical & Research Ethics committee (CREC) and NTW CREC in April 2010, and the subject recruitment in the Substance Abuse Clinic (SAC)s of CPH and NDH was proceeded

1.6 Revised study protocol

Revised sources of recruitment for both our original study and the collaborated study include patients attending the Substance Abuse Clinics (SACs) at PWH and other hospitals in the New Territories, namely North District Hospital (NDH) and Castle Peak Hospital (CPH); and clients with history of substance abuse from various non-governmental organizations (NGOs) in the New Territories.

Collection of data remained the same as our original study design except that subjects who completed the questionnaire and the urine collection would be paid with an honorarium. With the expanded sources of recruitment to the NTW, our study title is approved to be revised as "STUDY OF PATTERNS OF DRUGS OF ABUSE IN NEW TERRITORIES EAST AND WEST CLUSTER DRUG OF ABUSE CLINIC USING CONVENTIONAL AND NEW TECHNOLOGIES".

1.7 Recruitment

Subject recruitment started from July 2007 to March 2011. A total of 454 subjects were recruited. Among the 454 subjects, 275 subjects were male and 179 subjects were female with age ranged from 14 to 66 years.

2 Major Findings

2.1 Survey results

2.1.1 Demographic data

2.1.1.1 Age and sex distribution

| Age group (years) | Female | Male | Total | (N=454) |
|-------------------|--------|------|-------|---------|
| 10-19 | 30 | 20 | 50 | (11%) |
| 20-29 | 48 | 70 | 118 | (26%) |
| 30-39 | 49 | 110 | 159 | (35%) |
| 40-49 | 34 | 48 | 82 | (18.1%) |
| 50-59 | 16 | 25 | 41 | (9%) |
| 60-70 | 2 | 2 | 4 | (0.9%) |
| Grand Total | 179 | 275 | 454 | (100%) |

All subjects

*Age of the recruited subjects range from 14 to 66 years.

*Majority (72%) of the subjects is in the age groups below 40 years of age.

| Age group (years) | NTEC | (N=200) | NTWC | (N=254) | Total | (N=454) |
|-------------------|------|---------|------|---------|-------|---------|
| 10-19 | 34 | (17%) | 16 | (6.3%) | 50 | (11%) |
| 20-29 | 63 | (31.5%) | 55 | (21.7%) | 118 | (26%) |
| 30-39 | 70 | (35%) | 89 | (35%) | 159 | (35%) |
| 40-49 | 23 | (11.5%) | 59 | (23.2%) | 82 | (18.1%) |
| 50-59 | 9 | (4.5%) | 32 | (12.6%) | 41 | (9%) |
| 60-70 | 1 | (0.5%) | 3 | (1.2%) | 4 | (0.9%) |
| Grand Total | 200 | (100%) | 254 | (100%) | 454 | (100%) |

Comparing NTEC and NTWC

*Proportion of subjects with age of 40 years or above in NTWC (37%) is greater than that in NTEC (16.5%).

2.1.1.2 Marital Status

| Marital status | Female (N=179) | | Male | (N=275) | Total | (N=454) |
|----------------|----------------|---------|------|---------|-------|---------|
| Never married | 88 | (49.2%) | 151 | (54.9%) | 239 | (52.6%) |
| Co-habitant | 4 | (2.2%) | 2 | (0.7%) | 6 | (1.3%) |
| Married | 43 | (24%) | 78 | (28.4%) | 121 | (26.7%) |
| Divorced | 33 | (18.4%) | 28 | (10.2%) | 61 | (13.4%) |
| Separated | 3 | (1.7%) | 4 | (1.5%) | 7 | (1.5%) |
| Widowed | 2 | (1.1%) | 2 | (0.7%) | 4 | (0.9%) |
| Not specified | 6 | (3.4%) | 10 | (3.6%) | 16 | (3.5%) |
| Grand Total | 179 | (100%) | 275 | (100%) | 454 | (100%) |

*Majority of the subjects are single.

*A significant proportion of subjects is divorced.

2.1.1.3 Education attainment

| Education level | Female | Female (N=179) | | Male (N=275) | | (N=454) | | |
|---------------------|--------|----------------|-----|--------------|-----|---------|--|--|
| Primary education | 24 | (13.4%) | 43 | (15.6%) | 67 | (14.8%) | | |
| Secondary education | 150 | (83.8%) | 210 | (76.4%) | 360 | (79.3%) | | |
| Tertiary education | 3 | (1.7%) | 9 | (3.3%) | 12 | (2.6%) | | |
| Not specified | 2 | (1.1%) | 13 | (4.7%) | 15 | (3.3%) | | |
| Grand Total | 179 | (100%) | 275 | (100%) | 454 | (100%) | | |

Overall education level (by sex)

| Primary education up to | Female | Male | Total | (N=67) |
|-------------------------|--------|------|-------|---------|
| P.1 | | 1 | 1 | (1.5%) |
| P.2 | 3 | 3 | 6 | (9%) |
| P.3 | 2 | 4 | 6 | (9%) |
| P.4 | 3 | 2 | 5 | (7.5%) |
| P.5 | 3 | 7 | 10 | (14.9%) |
| P.6 | 13 | 26 | 39 | (58.2%) |
| Grand Total | 24 | 43 | 67 | (100%) |

| 1 | () / | | | |
|---------------------------|--------|------|-------|---------|
| Secondary education up to | Female | Male | Total | (N=360) |
| F.1 | 19 | 39 | 58 | (16.1%) |
| F.2 | 25 | 50 | 75 | (20.8%) |
| F.3 | 46 | 70 | 116 | (32.2%) |
| F.4 | 15 | 11 | 26 | (7.2%) |
| F.5 | 40 | 36 | 76 | (21.1%) |
| F.6 | 1 | 2 | 3 | (0.8%) |
| F.7 | 3 | 2 | 5 | (1.4%) |
| Not specified | 1 | 0 | 1 | (0.3%) |
| Grand Total | 150 | 210 | 360 | (100%) |
| | | | | |

Secondary education level (by sex)

| Tertiary education | |
|----------------------------------------|-------|
| Tertiary education | Total |
| Post-graduate (Master degree) | 1 |
| Undergraduate / Bachelor degree | 5 |
| Diploma | 1 |
| Not specified | 5 |
| Grand Total | 12 |

Overall education level (by age)

| Education level | Age group (Years) | | | | | | | |
|---------------------|-------------------|-------|-------|-------|-------|-------|-----|--|
| | 10-19 | 20-29 | 30-39 | 40-49 | 50-59 | 60-70 | | |
| Primary education | 1 | 2 | 7 | 27 | 27 | 3 | 67 | |
| Secondary education | 45 | 109 | 141 | 54 | 11 | | 360 | |
| Tertiary education | | 5 | 4 | 1 | 1 | 1 | 12 | |
| Not specified | 4 | 2 | 7 | | 2 | | 15 | |
| Grand Total | 50 | 118 | 159 | 82 | 41 | 4 | 454 | |

| Primary education | | Age group (Years) | | | | | |
|-------------------|-------|-------------------|-------|-------|-------|-------|----|
| up to | 10-19 | 20-29 | 30-39 | 40-49 | 50-59 | 60-70 | |
| P.1 | | | | | 1 | | 1 |
| P.2 | 1 | | | 1 | 4 | | 6 |
| P.3 | | | | 4 | 2 | | 6 |
| P.4 | | | | 3 | 1 | 1 | 5 |
| P.5 | | | 1 | 3 | 5 | 1 | 10 |
| P.6 | | 2 | 6 | 16 | 14 | 1 | 39 |
| Grand Total | 1 | 2 | 7 | 27 | 27 | 3 | 67 |

Primary education level (by age)

Secondary education level (by age)

| Secondary education | | Age group (Years) | | | | | Total |
|---------------------|-------|-------------------|-------|-------|-------|-------|-------|
| up to | 10-19 | 20-29 | 30-39 | 40-49 | 50-59 | 60-70 | |
| F.1 | 5 | 9 | 28 | 15 | 1 | | 58 |
| F.2 | 9 | 20 | 34 | 10 | 2 | | 75 |
| F.3 | 9 | 34 | 51 | 20 | 2 | | 116 |
| F.4 | 8 | 9 | 7 | 2 | | | 26 |
| F.5 | 14 | 32 | 20 | 6 | 4 | | 76 |
| F.6 | | 1 | 1 | | 1 | | 3 |
| F.7 | | 3 | | 1 | 1 | | 5 |
| Not specified | | 1 | | | | | 1 |
| Grand Total | 45 | 109 | 141 | 54 | 11 | 0 | 360 |

Tertiary level (by age)

| Tertiary education | | Age group (Years) | | | | | |
|--------------------------|-------|-------------------|-------|-------|-------|-------|----|
| up to | 10-19 | 20-29 | 30-39 | 40-49 | 50-59 | 60-70 | |
| Postgraudate (Master) | | | 1 | | | | 1 |
| Undergraduate / Bachelor | | 2 | 1 | 1 | 1 | | 5 |
| Diploma | | | 1 | | | | 1 |
| Not specified | | 3 | 1 | | | 1 | 5 |
| Grand Total | | 5 | 4 | 1 | 1 | 1 | 12 |

- *Majority (360 subjects, 79.3% of total) of the subjects attained secondary education level. Among these 360 subjects, 249 (69.1%) subjects attained junior secondary education level only.
- *These data is compatible with a general trend of increase in education level among the youngster population due to the provision of 9-year free education by the government since 1970s.

2.1.1.4 Current employment

| Employment status | Female | Male | Total | (N=454) |
|-------------------|--------|------|-------|---------|
| Unemployed | 121 | 166 | 287 | (63.2%) |
| Full-time | 23 | 56 | 79 | (17.4%) |
| Part-time | 11 | 33 | 44 | (9.7%) |
| Student | 9 | 12 | 21 | (4.6%) |
| Housewife | 13 | | 13 | (2.9%) |
| Retired | 1 | 3 | 4 | (0.9%) |
| Not specified | 1 | 5 | 6 | (1.3%) |
| Grand Total | 179 | 275 | 454 | (100%) |

Employment status (by sex)

Employment status (by age)

| Secondary education | | Age group (Years) | | | | | | | | | |
|---------------------|-------|-------------------|-------|-------|-------|-------|-----|--|--|--|--|
| up to | 10-19 | 20-29 | 30-39 | 40-49 | 50-59 | 60-70 | | | | | |
| Unemployed | 22 | 63 | 101 | 68 | 33 | | 287 | | | | |
| Full-time | 7 | 38 | 26 | 6 | 2 | | 79 | | | | |
| Part-time | 3 | 9 | 26 | 5 | | 1 | 44 | | | | |
| Student | 16 | 5 | | | | | 21 | | | | |
| Housewife | | 2 | 4 | 3 | 4 | | 13 | | | | |
| Retired | | | | | 1 | 3 | 4 | | | | |
| Not specified | 2 | 1 | 2 | | 1 | | 6 | | | | |
| Grand Total | 50 | 118 | 159 | 82 | 41 | 4 | 454 | | | | |

| Occupation reported among the 123 subjects currently under employment | | | | | | | | | | |
|-----------------------------------------------------------------------|----|---------------|----|--|--|--|--|--|--|--|
| Occupation | | | | | | | | | | |
| Customer service | 24 | Engineering | 3 | | | | | | | |
| Manual labour | 21 | Management | 1 | | | | | | | |
| Food/Catering | 11 | Banking | 1 | | | | | | | |
| Transport/Driver | 10 | Others | 5 | | | | | | | |
| Healthcare | 4 | Not specified | 43 | | | | | | | |
| Total: 123 subjects | | | | | | | | | | |

Occupation reported among the 123 subjects currently under employment

*A total of 287 (63.2%) subjects are currently unemployed. This high overall unemployment rate among our study population is 18-fold greater than the unemployment rate of the general population. [Source: According to the latest labor force statistics released by the Census and Statistics Department (C&SD) of the Hong Kong Special Administrative Region Government, Hong Kong's seasonally adjusted unemployment rate was provisional 3.5% in February - April 2011.]

*High unemployment rates among the younger age groups, 53.4% and 63.5% in the 20-29 years and 30-39 years age groups respectively, as shown by our data, are particularly of significance as people of these age groups are supposed to be the major workforce of the society. In the 40-49 years and 50-59 years age groups, the unemployment rates are even higher to over 80% in the corresponding age group. (Note: Percentages are calculated based on the subjects of the corresponding age group.)

2.1.1.5 Monthly personal income and expenditure on drugs

| Employment status | Female | Male | Total | (N=454) |
|---------------------|--------|------|-------|---------|
| No income | 142 | 172 | 314 | (69.2%) |
| < \$1,000 | 5 | 4 | 9 | (2%) |
| \$1,000 to \$2,999 | 4 | 10 | 14 | (3.1%) |
| \$3,000 to \$4,999 | 7 | 15 | 22 | (4.8%) |
| \$5,000 to 6,999 | 8 | 11 | 19 | (4.2%) |
| \$7,000 to \$8,999 | 3 | 20 | 23 | (5.1%) |
| \$9,000 to \$10,999 | 4 | 13 | 17 | (3.7%) |
| >\$11,000 | 5 | 23 | 28 | (6.2%) |
| Not specified | 1 | 7 | 8 | (1.8%) |
| Grand Total | 179 | 275 | 454 | (100%) |

Monthly personal income (by sex)

*With the high unemployment rate among our study population, majority of subjects have no monthly personal income. Among the 314 subjects who have no income, 209 subjects (66.6% of 314 subjects) reported to be receiving financial assistance from the government.

| Expenditure | Female | Male | Total | (N=454) |
|------------------|--------|------|-------|---------|
| Nil | 80 | 132 | 212 | (46.7%) |
| <\$100 | 7 | 8 | 15 | (3.3%) |
| \$100 to \$299 | 15 | 20 | 35 | (7.7%) |
| \$300 to \$499 | 12 | 15 | 27 | (5.9%) |
| \$500 to \$699 | 9 | 8 | 17 | (3.7%) |
| \$700 to \$899 | 5 | 4 | 9 | (2%) |
| \$900 to \$1,099 | 12 | 6 | 18 | (4%) |
| >\$1,100 | 38 | 75 | 113 | (24.9%) |
| Not specified | 1 | 7 | 8 | (1.8%) |
| Grand Total | 179 | 275 | 454 | (100%) |

Monthly expenditure on drugs (by sex) \geq

| Expenditure | No income | | Monthly incom | e | Not |
|------------------|-------------|------------|---------------|-----------|-----------|
| | | <\$5000 | \$5000 to | >\$11,000 | specified |
| | | | \$10,999 | | |
| | (N=314) | (N=45) | (N=59) | (N=28) | (N=8) |
| Nil | 150 (47.8%) | 24 (53.3%) | 28 (47.5%) | 8 (28.6%) | 2 (25%) |
| <\$100 | 11 (3.5%) | 2 (4.4%) | 1 (1.7%) | 1 (3.6%) | |
| \$100 to \$299 | 24 (7.6%) | 7 (15.6%) | 2 (3.4%) | 2 (7.1%) | |
| \$300 to \$499 | 20 (6.4%) | 4 (8.9%) | 2 (3.4%) | 1 (3.6%) | |
| \$500 to \$699 | 11 (3.5%) | 2 (4.4%) | 3 (5.1%) | 1 (3.6%) | |
| \$700 to \$899 | 8 (2.5%) | 1 (2.2%) | | | |
| \$900 to \$1,099 | 16 (5.1%) | 1 (2.2%) | | 1 (3.6%) | |
| >\$1,100 | 71 (22.6%) | 4 (8.9%) | 23 (39%) | 14 (50%) | 1 (12.5%) |
| Not specified | 3 (1%) | | | | 5 (62.5%) |
| Grand Total | 314 (100%) | 45 (100%) | 59 (100%) | 28 (100%) | 8 (100%) |

Monthly expenditure on drugs by income

*As unemployment could be a source of stress to the drug abusers that promotes their continuation of substance abuse behavior, 161 (51.3%) of 314 subjects who have no income reported to have monthly expenditure on drugs ranging from <\$100 to >\$1,100. Among these 161 subjects, 111 of them were receiving financial assistance from government.

*However, for those who are employed and have monthly income >\$5,000, 42.5% (37 of 87) of subjects reported to have monthly expenditure on drugs of >\$1,100.

2.1.2 Information of previous drug use

2.1.2.1 Age first started and duration of drug abuse

| Age group | Female | | Male | | Total | |
|---------------|--------|---------|------|---------|-------|---------|
| | N | % | N | % | N | % |
| 10-14 | 38 | (21.2%) | 52 | (18.9%) | 90 | (19.8%) |
| 15-19 | 74 | (41.3%) | 127 | (46.2%) | 201 | (44.3%) |
| 20-24 | 24 | (13.4%) | 47 | (17.1%) | 71 | (15.6%) |
| 25-29 | 10 | (5.6%) | 15 | (5.5%) | 25 | (5.5%) |
| 30-34 | 11 | (6.1%) | 12 | (4.4%) | 23 | (5.1%) |
| 35-39 | 9 | (5.0%) | 3 | (1.1%) | 12 | (2.6%) |
| 40-44 | 6 | (3.4%) | 1 | (0.4%) | 7 | (1.5%) |
| 45-49 | 3 | (1.7%) | 1 | (0.4%) | 4 | (0.9%) |
| 50-54 | 1 | (0.6%) | 4 | (1.5%) | 5 | (1.1%) |
| 55-60 | 1 | (0.6%) | 1 | (0.4%) | 2 | (0.4%) |
| Not specified | 2 | (1.1%) | 12 | (4.4%) | 14 | (3.1%) |
| Grand Total | 179 | (100%) | 275 | (100%) | 454 | (100%) |

| Age first started drug abuse (by sex) | (by sex) | drug abuse | Age first started | \triangleright |
|---------------------------------------|----------|------------|-------------------|------------------|
|---------------------------------------|----------|------------|-------------------|------------------|

*Age first started drug abuse in our study population was reported to be ranged from 10 to 57 years.

*Pattern of age of first started drug abuse is quite similar in both sex with majority of subjects first started drug abuse at age between 15 and 19 years and a significant proportion of subjects reported an even earlier age of starting drug abuse between 10 to 14 years. This finding is coherent with the findings of other studies that drug of abuse problem is now a major issue among our youngsters.

| Duration | | | | | | Age grou | o (Year | s) | | | | | | |
|---------------|------|--------|-------|---------|-------|----------|---------|---------|------|---------|-----|--------|-------|---------|
| Duration | 10-1 | 19 | 20-29 | Э | 30-39 | 9 | 40-4 | 9 | 50-5 | 9 | 60- | -69 | Total | |
| (Years) | Ν | % | N | % | N | % | N | % | Ν | % | Ν | % | N | % |
| < 1 year | 1 | (2%) | 1 | (0.8%) | | | | | | | | | 2 | (0.4%) |
| 1-5 years | 44 | (88%) | 28 | (23.7%) | 5 | (3.1%) | 6 | (7.3%) | 5 | (12.2%) | | | 88 | (19.4%) |
| 6-10 years | 4 | (8%) | 56 | (47.5%) | 22 | (13.8%) | 8 | (9.8%) | 3 | (7.3%) | 2 | (50%) | 95 | (20.9%) |
| 11-15 years | | | 23 | (19.5%) | 32 | (20.1%) | 7 | (8.5%) | 2 | (4.9%) | | | 64 | (14.1%) |
| 16-20 years | | | 6 | (5.1%) | 74 | (46.5%) | 15 | (18.3%) | 6 | (14.6%) | | | 101 | (22.2%) |
| 21-25 years | | | | | 18 | (11.3%) | 11 | (13.4%) | 1 | (2.4%) | 1 | (25%) | 31 | (6.8%) |
| 26-30 years | | | | | 1 | (0.6%) | 30 | (36.6%) | 6 | (14.6%) | | | 37 | (8.1%) |
| 31-35 years | | | | | | | 5 | (6.1%) | 8 | (19.5%) | 1 | (25%) | 14 | (3.1%) |
| 36-40 years | | | | | | | | | 6 | (14.6%) | | | 6 | (1.3%) |
| 41-45 years | | | | | | | | | 2 | (4.9%) | | | 2 | (0.4%) |
| Not specified | 1 | (2%) | 4 | (3.4%) | 7 | (4.4%) | | | 2 | (4.9%) | | | 14 | (3.1%) |
| Grand Total | 50 | (100%) | 118 | (100%) | 159 | (100%) | 82 | (100%) | 41 | (100%) | 4 | (100%) | 454 | (100%) |

Duration of drug abuse

*Duration of drug abuse was calculated by subtracting the age first started drug abuse from the current age reported.

*Percentage is calculated based on the number (N) of each age group and the total number of recruited subjects.

*For those 59 subjects who have drug abuse for more than 25 years, 46 (78%) of them still have positive drug detection in their urine samples. Opiates (24 samples), hypnotics (17 samples), benzodiazepines (11 samples), cough medicine (11 samples), amphetamines (4 samples), cannabis (1 sample) and barbiturate (1 sample) were detected. (Note: One or more drug items were detected in individual urine sample.)

| Duration of follow up | Female | | Male | | Total | |
|-----------------------|--------|---------|------|---------|-------|---------|
| (Years) | Ν | % | Ν | % | Ν | % |
| New case | 23 | (12.8%) | 35 | (12.7%) | 58 | (12.8%) |
| < 1 year | 30 | (16.8%) | 32 | (11.6%) | 62 | (13.7%) |
| 1-5 years | 55 | (30.7%) | 87 | (31.6%) | 142 | (31.3%) |
| 6-10 years | 25 | (14%) | 42 | (15.3%) | 67 | (14.8%) |
| > 10 years | 4 | (2.2%) | 11 | (4%) | 15 | (3.3%) |
| Not specified | 42 | (23.5%) | 68 | (24.7%) | 110 | (24.2%) |
| Grand Total | 179 | (100%) | 275 | (100%) | 454 | (100%) |

Duration of follow up in SAC

*Note: SACs in PWH and CPH started service since 1995.

*For old cases, the reported duration of follow up in SAC ranged from 2 weeks to 16 years.

2.1.2.2 Groups of drug items ever tried

| Number of groups | Femal | Female | | | Total | Total | | |
|------------------|-------|---------|-----|---------|-------|---------|--|--|
| of drug items | Ν | % | Ν | N % | | % | | |
| 1 | 48 | (26.8%) | 55 | (20.0%) | 103 | (22.7%) | | |
| 2 | 23 | (12.8%) | 36 | (13.1%) | 59 | (13.0%) | | |
| 3 | 18 | (10.1%) | 41 | (14.9%) | 59 | (13.0%) | | |
| 4 | 20 | (11.2%) | 34 | (12.4%) | 54 | (11.9%) | | |
| 5 | 28 | (15.6%) | 39 | (14.2%) | 67 | (14.8%) | | |
| 6 | 12 | (6.7%) | 17 | (6.2%) | 29 | (6.4%) | | |
| 7 | 12 | (6.7%) | 18 | (6.5%) | 30 | (6.6%) | | |
| 8 | 13 | (7.3%) | 12 | (4.4%) | 25 | (5.5%) | | |
| 9 | 4 | (2.2%) | 13 | (4.7%) | 17 | (3.7%) | | |
| 10 | 1 | (0.6%) | 5 | (1.8%) | 6 | (1.3%) | | |
| Not specified | | | 5 | (1.8%) | 5 | (1.1%) | | |
| Grand Total | 179 | (100%) | 275 | (100%) | 454 | (100%) | | |

> Number of groups of drug items ever tried (by sex)

*Note: Drug items ever tried by individual subject since his/her first start to the time of the survey were counted.

*Number of drug items ever tried by individual subject ranged from 1 to 10.

- *Percentage is calculated based on the number (N) of corresponding sex and the total number of recruited subjects.
- *Only 22.7% of the 454 subjects reported to be single drug users all along while 76.2% reported to have tried more than one group of drugs in the past.
- *Both sex showed similar pattern.

| Number of groups of | | | Total | | | | | |
|---------------------|-------|-------|-------|-------|-------|-------|-----|---------|
| drug items | 10-19 | 20-29 | 30-39 | 40-49 | 50-59 | 60-70 | | (N=454) |
| 1 | 11 | 24 | 26 | 23 | 16 | 3 | 103 | (22.7%) |
| 2 | 8 | 12 | 19 | 14 | 6 | | 59 | (13.0%) |
| 3 | 7 | 14 | 20 | 14 | 4 | | 59 | (13.0%) |
| 4 | 8 | 17 | 21 | 3 | 5 | | 54 | (11.9%) |
| 5 | 10 | 21 | 20 | 11 | 4 | 1 | 67 | (14.8%) |
| 6 | 3 | 10 | 9 | 6 | 1 | | 29 | (6.4%) |
| 7 | 2 | 8 | 15 | 4 | 1 | | 30 | (6.6%) |
| 8 | | 7 | 14 | 2 | 2 | | 25 | (5.5%) |
| 9 | | 3 | 10 | 3 | 1 | | 17 | (3.7%) |
| 10 | | 1 | 3 | 2 | | | 6 | (1.3%) |
| Not specified | 1 | 1 | 2 | | 1 | | 5 | (1.1%) |
| Grand Total | 50 | 118 | 159 | 82 | 41 | 4 | 454 | (100%) |

> Number of groups of drug items ever tried (by age)

*Proportion of multiple drug users is the greatest among the 30-39 years age group. Of which, 57.9% subjects reported to have tried 4 or more drugs.

Summary of group of drug item ever tried (by sex)

| Group of drug item | Femal | Female | | Male | | |
|-------------------------|-------|---------|------|---------|------|-------|
| | (N=17 | (N=179) | | (N=275) | | 54) |
| | | % | | % | | % |
| Ketamine | 102 | 57.0% | 148 | 53.8% | 250 | 55.1% |
| Cannabis | 82 | 45.8% | 154 | 56.0% | 236 | 52.0% |
| Amphetamines | 98 | 54.7% | 137 | 49.8% | 235 | 51.8% |
| Hypnotics | 86 | 48.0% | 96 | 34.9% | 182 | 40.1% |
| Opiates | 61 | 34.1% | 111 | 40.4% | 172 | 37.9% |
| Benzodiazepines | 69 | 38.5% | 101 | 36.7% | 170 | 37.4% |
| MDMA (Ecstasy) | 61 | 34.1% | 106 | 38.5% | 167 | 36.8% |
| Cough medicine | 42 | 23.5% | 115 | 41.8% | 157 | 34.6% |
| Cocaine | 59 | 33.0% | 66 | 24.0% | 125 | 27.5% |
| Mandrax | 14 | 7.8% | 27 | 9.8% | 41 | 9.0% |
| Others | 2 | 1.1% | 5 | 1.8% | 7 | 1.5% |
| Thinner/Organic solvent | 1 | 0.6% | 6 | 2.2% | 7 | 1.5% |
| Grand Total | 677 | | 1072 | | 1749 | |

*Data shown in descending order of the overall popularity.

- *Note: Groups of drug items ever tried by individual subject since his/her first start to the time of the survey are counted. An individual subject can reported one or more groups of drug items tried. Figures are presented as the number of counts reported and the percentage is calculated based on the number (N) of corresponding sex and the total number of recruited subjects.
- *Ketamine was the most popular group of drug that was ever tried by our study population (55.1% of 454 subjects).

| Group of drug item | | Age group (Years) | | | | | | Total | | | | | | |
|-------------------------|-------|-------------------|-------|---------|--------|---------|-------|---------|-------|---------|----|-------|---------|---------|
| | 10-19 | | 20-29 | | 30-39 | | 40-49 | | 50-59 | | 60 | -70 | | |
| | (N=50 |)) | (N=11 | 8) | (N=159 |) | (N=82 |) | (N=41 |) | (N | =4) | (N=454) | |
| Ketamine | 43 | (86%) | 101 | (85.6%) | 83 | (52.2%) | 16 | (19.5%) | 7 | (17.1%) | | | 250 | (55.1%) |
| Cannabis | 19 | (38%) | 67 | (56.8%) | 98 | (61.6%) | 38 | (46.3%) | 13 | (31.7%) | 1 | (25%) | 236 | (52%) |
| Amphetamines | 29 | (58%) | 63 | (53.4%) | 99 | (62.3%) | 35 | (42.7%) | 9 | (22%) | | | 235 | (51.8%) |
| Hypnotics | 4 | (8%) | 29 | (24.6%) | 67 | (42.1%) | 50 | (61%) | 29 | (70.7%) | 3 | (75%) | 182 | (40.1%) |
| Opiates | 3 | (6%) | 18 | (15.3%) | 81 | (50.9%) | 44 | (53.7%) | 24 | (58.5%) | 2 | (50%) | 172 | (37.9%) |
| Benzodiazepines | 10 | (20%) | 41 | (34.7%) | 59 | (37.1%) | 40 | (48.8%) | 19 | (46.3%) | 1 | (25%) | 170 | (37.4%) |
| MDMA (Ecstasy) | 21 | (42%) | 66 | (55.9%) | 61 | (38.4%) | 14 | (17.1%) | 5 | (12.2%) | | | 167 | (36.8%) |
| Cough medicine | 6 | (12%) | 30 | (25.4%) | 89 | (56%) | 28 | (34.1%) | 4 | (9.8%) | | | 157 | (34.6%) |
| Cocaine | 21 | (42%) | 48 | (40.7%) | 43 | (27%) | 10 | (12.2%) | 2 | (4.9%) | 1 | (25%) | 125 | (27.5%) |
| Mandrax | 5 | (10%) | 8 | (6.8%) | 15 | (9.4%) | 7 | (8.5%) | 6 | (14.6%) | | | 41 | (9%) |
| Thinner/Organic solvent | | | 4 | (3.4%) | 1 | (0.6%) | 2 | (2.4%) | | | | | 7 | (1.5%) |
| Others | 1 | (2%) | | | 4 | (2.5%) | 2 | (2.4%) | | | | | 7 | (1.5%) |
| Grand Total | 162 | | 475 | | 700 | | 286 | | 118 | | 8 | | 1749 | |

Summary of group of drug item ever tried by age

*Data shown in descending order of the overall popularity.

- *Note: Groups of drug items ever tried by individual subject from his/her first attempt to the time of the survey are counted. An individual subject can reported one or more groups of drug items tried. Figures are presented as the number of counts reported. Percentage is calculated based on the total number (N) of subjects of the corresponding age groups.
- *Comparing the pattern of drug ever tried across different age groups, different popularity of drug use was shown.
- *In 10-29 year group, ketamine was the most popular drug.
- *In 30-39 year group, amphetamine was the most popular drug.

*In 40-70 year group, hypnotics was the most popular drug.

- *Overall, ketamine, cannabis and amphetamines were the three most common drugs being abused in our study population.
- *The reported usage of cough medicine was relatively low except that in the 30-39 years age group comparing with the high frequency of detection in the urine samples.

| Reason of drug abuse | Female (N=179) | | Male (N=275 | Male (N=275) | | 1) |
|--------------------------------------|-------------------|---------|----------------|-----------------|-----|---------|
| | | % | | % | | % |
| Peer influence | 70 | (39.1%) | 131 | (47.6%) | 201 | (44.3%) |
| Recreational | 52 | (29.1%) | 81 | (29.5%) | 133 | (29.3%) |
| Experimental | 25 | (14.0%) | 52 | (18.9%) | 77 | (17.0%) |
| Insomnia | 28 | (15.6%) | 7 | (2.5%) | 35 | (7.7%) |
| Dependence | 13 | (7.3%) | 9 | (3.3%) | 22 | (4.8%) |
| Unhappiness/Stress | 12 | (6.7%) | 8 | (2.9%) | 20 | (4.4%) |
| Family/Marriage/Relationship Affairs | 11 | (6.1%) | 2 | (0.7%) | 13 | (2.9%) |
| Cough/Asthma | 1 | (0.6%) | 4 | (1.5%) | 5 | (1.1%) |
| Weight reduction | 2 | (1.1%) | | | 2 | (0.4%) |
| Not specified | 4 | (2.2%) | 4 | (1.5%) | 8 | (1.8%) |
| Grand Total | 218 | | 298 | | 516 | |

2.1.2.3 Reason of drug abuse

*Note: Subject can report one or more reasons of drug abuse. Figures are presented as the number of counts reported. Percentage is calculated based on the total number (N) of subjects of the corresponding sex.

*Peer influence and recreational are the two major reasons of drug abuse reported in our study population.

| Presence of complications | Female | | Male | | Tota | l |
|---------------------------|--------|---------|------|---------|------|---------|
| | Ν | % | Ν | % | Ν | % |
| Yes | 129 | (72.1%) | 191 | (69.5%) | 320 | (70.5%) |
| No | 44 | (24.6%) | 66 | (24%) | 110 | (24.2%) |
| Not specified | 6 | (3.4%) | 18 | (6.5%) | 24 | (5.3%) |
| Grand Total | 179 | (100%) | 275 | (100%) | 454 | (100%) |

2.1.2.4 Subjective complications of drug abuse

*Most (70.5%) of the subjects reported to have experienced complications of drug abuse subjectively.

*Physical complications frequently that were most reported include tiredness/weakness, hand tremor, dental caries, urinary frequency/ketamine-induced cystitis, loss of appetite, withdrawal symptoms, epigastric pain and hepatitis. Serious physical complications such as infective endocarditis, heart failure, deep vein thrombosis, loss of consciousness due to drug overdose were also reported.

*Psychiatric complications that were most frequently reported include impaired memory, depression, psychosis, hallucination, anxiety, irritability, insomnia, slow response and suicidal idea/attempt.

2.1.3 Information of last drug use

2.1.3.1 Estimated time of last use

| When was last drug use? | Female | Male | Total | (N=454) |
|-----------------------------------------------|--------|------|-------|---------|
| Less than 1 day | 41 | 55 | 96 | (21.1%) |
| 1 day to less than or equal to 1 week | 49 | 82 | 131 | (28.9%) |
| Above 1 week to less than or equal to 1 month | 27 | 31 | 58 | (12.8%) |
| Above 1 month to less than or equal to 1 year | 39 | 69 | 108 | (23.8%) |
| Above 1 year | 21 | 23 | 44 | (9.7%) |
| Quitted for unspecified period of time | | 2 | 2 | (0.4%) |
| Not specified | 2 | 13 | 15 | (3.3%) |
| Grand Total | 179 | 275 | 454 | (100%) |

*A total of 227 (50%) subjects reported his/her last drug use to be within 1 week's time with regard to the time of the survey and urine collection, in which, 96 (21.1%) subjects admitted that they had active drug use even less than 24 hours before the clinic follow up.

*A total of 152 (33.5%) subjects reported a drug-free period of above 1 month, in which, only 44 (9.7%) subjects reported his/her last drug use to be above 1 year of time.

2.1.3.2 Drug items of last use

| Number of drug item last used | Female | Male | Total | (N=454) |
|-------------------------------|--------|------|-------|---------|
| 1 | 156 | 241 | 397 | 87.4% |
| 2 | 15 | 19 | 34 | 7.5% |
| 3 | 3 | 1 | 4 | 0.9% |
| 4 | 3 | 1 | 4 | 0.9% |
| Not specified | 2 | 13 | 15 | 3.3% |
| Grand Total | 179 | 275 | 454 | (100%) |

Number of drug item last used

*Most (87.4%) subjects reported using one drug item only.

| Group of drug item last used | Fema | | Mal | | Total | |
|-------------------------------|---------|---------|---------|---------|---------|---------|
| | (N=179) | | (N=275) | | (N=454) | |
| | | % | | % | | % |
| Ketamine | 44 | (24.6%) | 69 | (25.1%) | 113 | (24.9%) |
| Hypnotics | 48 | (26.8%) | 35 | (12.7%) | 83 | (18.3%) |
| Opiates | 29 | (16.2%) | 53 | (19.3%) | 82 | (18.1%) |
| Cough medicine | 14 | (7.8%) | 64 | (23.3%) | 78 | (17.2%) |
| Amphetamines | 43 | (24%) | 29 | (10.5%) | 72 | (15.9%) |
| Benzodiazepines | 15 | (8.4%) | 10 | (3.6%) | 25 | (5.5%) |
| Cannabis | 2 | (1.1%) | 12 | (4.4%) | 14 | (3.1%) |
| Cocaine | 7 | (3.9%) | 6 | (2.2%) | 13 | (2.9%) |
| MDMA (Ecstasy) | 2 | (1.1%) | 4 | (1.5%) | 6 | (1.3%) |
| Mandrax | 3 | (1.7%) | | | 3 | (0.7%) |
| Others (e.g. LSD, analgesics) | | | 2 | (0.7%) | 2 | (0.4%) |
| Not specified | 2 | (1.1%) | 13 | (4.7%) | 15 | (3.3%) |
| Grand Total | 209 | | 297 | | 506 | |

Summary of group of drug item last used by sex (Data shown in descending order of the overall popularity)

*Note: Individual subject could report more than one drug item during his/her last use. Figures are presented as the number of counts reported. Percentage is calculated based on the total number (N) of subjects of the corresponding sex.

- *Hypnotics reported include zopiclone, zolpidem and unknown hypnotics bought over-the-counter.
- *Benzodiazepines reported include diazepam, midazolam, nimetazepam and others.
- *Ketamine, hypnotics, opiates (heroin), cough medicine and amphetamines are the five most popular groups of drug items that were reported by our study subjects during their last drug use.
- *Cough medicine is the second most popular group of drug item reported in male subjects while its use is less popular in female subjects.

2.1.3.3 Source of drugs of abuse

| | 5 | 18 Districts of | Hong Kong | | | |
|---------------------------|-----|-----------------|-----------|-------------------|---|--|
| 18 Districts of Hong Kong | | | | | | |
| New Territories | | Kowloon | | Hong Kong Island | | |
| (Subtotal: 298) | | (Subtotal: 116) | | (Subtotal: 7) | | |
| Yuen Long | 121 | Yau Tsim Mong | 88 | Central & Western | 3 | |
| Tuen Mun | 86 | Kwun Tong | 14 | Eastern | 2 | |
| Shatin | 43 | Sham Shui Po | 9 | Wan Chai | 1 | |
| North | 28 | Kowloon City | 4 | Southern | 1 | |
| Tai Po | 12 | Wong Tai Sin | 1 | | | |
| Tsuen Wan | 4 | | | | | |
| Islands | 2 | | | | | |
| Sai Kung | 2 | | | | | |
| Kwai Ching | 0 | | | | | |
| (Grand Total: 421) | | | | | | |

Source of drugs from Hong Kong region

| Source of drugs outside Ho | ng Kong region |
|--------------------------------|----------------|
| Outside Hong Kong | |
| Shenzhen | 16 |
| Mainland China (not specified) | 2 |
| (Grand Total: 18) | |

*Note: Individual subject could report more than one district or region for the source of drug obtained. Some subjects did not report the source of drugs. Figures are presented as the number of counts reported.

*Most of the study subjects obtained drugs in Hong Kong, particularly in Yuen Long, Tuen Mun and Yau Tsim Mong districts.

Channels to obtain drugs

| Channels | Total | (N = 454) |
|---------------|-------|-----------|
| Friends | 199 | (43.8%) |
| Pharmacy | 142 | (31.3%) |
| Others | 85 | (18.7%) |
| Not specified | 33 | (7.3%) |
| Grand Total | 459 | (100%) |

*Note: Individual subject could report more than one channel. Figures are presented as the number of counts reported. Percentage is calculated based on the total number (N) of all subjects.

*Friends are reported as common sources of ketamine and amphetamines

* Pharmacy is reported as a common source of hypnotics and cough medicine.

2.1.3.4 Location of drug use

| Location of drug use | Femal | e | Male | | Total | |
|---------------------------|-------|---------|------|---------|-------|---------|
| Location of urug use | (N=17 | (N=179) | | (N=275) | |) |
| | | % | | % | | % |
| Home | 117 | (65.4%) | 124 | (45.1%) | 241 | (53.1%) |
| Friend's Apartment | 39 | (21.8%) | 34 | (12.4%) | 73 | (16.1%) |
| Street/Public toilet/Park | 11 | (6.1%) | 60 | (21.8%) | 71 | (15.6%) |
| Disco | 5 | (2.8%) | 22 | (8%) | 27 | (5.9%) |
| Pharmacy | 2 | (1.1%) | 24 | (8.7%) | 26 | (5.7%) |
| Video Game Centre | | | 6 | (2.2%) | 6 | (1.3%) |
| Karaoke | 3 | (1.7%) | 3 | (1.1%) | 6 | (1.3%) |
| Bar | 2 | (1.1%) | 3 | (1.1%) | 5 | (1.1%) |
| Hotel | | | 1 | (0.4%) | 1 | (0.2%) |
| Workplace | | | 1 | (0.4%) | 1 | (0.2%) |
| Not specified | 3 | (1.7%) | 10 | (3.6%) | 13 | (2.9%) |
| Grand Total | 182 | | 288 | | 470 | |

*Note: Individual subject could report more than one location of drug use. Figures are presented as the number of counts reported. Percentage is calculated based on the total number (N) of subjects of the corresponding sex.

*Majority of subjects reported to use drugs at home or friend's apartment.

*Location of drug use reported in female and male subjects has slightly different patterns.

2.1.4 Other data

2.1.4.1 Current drug history

| \triangleright | Currently on therapeutic drugs |
|------------------|--------------------------------|
| | |

| On therapeutic drug therapy | Female | Male | Total | (N=454) |
|-----------------------------|--------|------|-------|---------|
| Yes | 128 | 170 | 298 | (65.6%) |
| No | 50 | 96 | 146 | (32.2%) |
| Not specified | 1 | 9 | 10 | (2.2%) |
| Grand Total | 179 | 275 | 454 | (100%) |

*Note: Therapeutic drug items used by the study subjects are mainly psychiatric medications prescribed from drug of abuse clinic. However, detail description of individual's therapeutic drug regime is unavailable.

2.1.4.2 Smoking and drinking history

Smoking history

| Smoking history | Female | Male | Total | (N=454) |
|-----------------|--------|------|-------|---------|
| Smoker | 137 | 245 | 382 | (84.1%) |
| Ex-smoker | 7 | 10 | 17 | (3.7%) |
| Non-smoker | 30 | 12 | 42 | (9.3%) |
| Not specified | 5 | 8 | 13 | (2.9%) |
| Grand Total | 179 | 275 | 454 | (100%) |

Drinking history

| Smoking history | Female | Male | Total | (N=454) |
|-----------------|--------|------|-------|---------|
| Drinker | 56 | 121 | 177 | (39%) |
| Ex-drinker | 28 | 30 | 58 | (12.8%) |
| Non-drinker | 91 | 112 | 203 | (44.7%) |
| Not specified | 4 | 12 | 16 | (3.5%) |
| Grand Total | 179 | 275 | 454 | (100%) |

*Most (84.1%) study subjects reported to be smokers.

*Overall proportion of drinkers and non-drinkers among our study population is similar.

2.1.4.3 Forensic history

Forensic history by sex

| Forensic history | Female | Female | | Male | | |
|------------------|--------|---------|-----|---------|-----|---------|
| | Ν | % | Ν | % | Ν | % |
| Yes | 79 | (44.1%) | 197 | (71.6%) | 276 | (60.8%) |
| No | 96 | (53.6%) | 68 | (24.7%) | 164 | (36.1%) |
| Not specified | 4 | (2.2%) | 10 | (3.6%) | 14 | (3.1%) |
| Grand Total | 179 | (100%) | 275 | (100%) | 454 | (100%) |

*A total of 276 (60.8%) study subjects reported to have previous forensic history.

*Proportion of male subjects reported to have forensic history is higher than that of female subjects.

| Summary of forensic history | |
|-----------------------------------|------------------|
| Drug-related forensic history | Subtotal: 178 |
| Drug possession | 129 |
| Drug trafficking | 36 |
| Drug abusing | 13 |
| Non-drug-related forensic history | Subtotal: 153 |
| Theft | 64 |
| Assault | 45 |
| Robbery | 24 |
| Others | 20 |
| Not specified | Subtotal: 52 |
| | Grand Total: 383 |
| | |

Summary of forensic history

*Note: Individual subject could report more than one item. Others include money laundering, uttering/forgery, blackmail, attacking police, possession of offensive weapon, destruction of property, drug-driving, driving without license, child abuse and other unspecified crime.

2.2 Results of urine analysis

2.2.1 Overall results

2.2.1.1 Overview of urine analysis

A total of 454 urine samples were collected from July 2007 to March 2011.

All urine samples were collected using the temperature-sensitive urine bottles and were aliquoted for urine creatinine measurement to ensure the validity of the urine specimens. Twenty-one (4.6%) of the 454 urine samples were found to have urine creatinine level below 1.8 mmol/L that were regarded as abnormally diluted urine, yet drugs/metabolites were still detectable in these urine samples.

All urine samples were subjected to a comprehensive drug screening by UPLC-TOF-MS (Waters UPLC-LCT Premier XE) and a panel of drug of abuse screening using Immunoassay (Abbott AxSym® System). Selected urine samples were also analyzed by HPLC-UV (REMEDi HS drug profiling system) for the comparison study.

➢ UPLC-TOF-MS

UPLC-TOF-MS can be used for broad toxicological screening based on the exact mass determination with the advantage of generation of theoretical exact mass libraries from the molecular formula for drug identification without the need of reference materials. The UPLC-TOF-MS method used in our study has been validated and published in a peer review journal. Utility and overall performance of the UPLC-TOF-MS method were assessed by the analysis of 30 authentic urine samples by both UPLC-TOF-MS (Waters UPLC-LCT Premier XE) and HPLC-UV (REMEDi HS drug profiling system). The study showed that the UPLC-TOF-MS detected twice as many as drugs as the HPLC-UV. The authentic urine samples have been sent to Sweden and UK for further comparison using other conventional screening techniques including, high performance liquid chromatography-diode array detector (HPLC-DAD), liquid chromatography tandem mass spectrometry (LC-MS/MS) and gas chromatography mass spectrometry (GC-MS). These additional methods confirmed no false positive detected by the UPLC-TOF-MS method. Within-instrument precisions for retention time and signal responses (peak area) were satisfactory; coefficients of variation (CV) were 0.29% and 11.3%, respectively. Lower detection

limit ranged from 1 to 125 ng/mL. The detail of the materials, instrumentation, experimental studies and validation data were published. [HK Lee *et al*. Anal Chim Acta (2009) 649:80-90]

Advanced software systems, ChromaLynxTM and QuanLynxTM (Waters), were used for broad-spectrum drug screening and targeted analysis respectively for detection of drugs by the UPLC-TOF-MS system. The software provides an automated identification of component peaks using spectral deconvolution techniques followed by comparison of the underlying mass spectra with the entries in the drug libraries containing more than 300 common drugs and metabolites.

Positive detection of a drug or metabolite by UPLC-TOF-MS was defined by a combination of criteria, including an average spectral match factor (which was based on the nominal mass spectra acquired under low and high fragmentation conditions of each candidate comparing against the respective entries in the libraries) \geq 500, retention time within 10% of expected, accurate mass measurement < 5 parts per million (ppm) and compatible elemental composition and isotopic pattern (I-fit).

HPLC-UV

The REMEDi HS drug profiling system is a broad spectrum drug identification system using liquid chromatography with on-line sample cleanup and isocratic multicolumn separation with full-scan UV detection.

For sample processing, the samples are diluted with an internal standard mixture and centrifuged. The internal standards are used to monitor the chromatographic behavior of the separation cartridges. Upon injection, the prepared sample is combined with a buffer and passed through four cartridges. All reagents used with the Remedi were supplied by Bio-Rad; however, the composition of the mobile phase and other reagents and the exact characteristics of the stationary phase of the cartridges were not disclosed.

Drug identification is performed by a multi-wavelength UV detector coupled with a sophisticated computer algorithm. As each drug enters the detector, a UV scan from 200 nm to 300 nm is made. Sample spectra are then automatically compared with the library of known drug spectra stored in memory. This, in conjunction with chromatographic data, results in the identification of the drug.

Immunoassay

The Abbott AxSym® System utilizes Fluorescence Polarization Immunoassay (FPIA) technology for detection of drugs and metabolites in urine specimen. A panel of immunoassays covering the seven common groups of drug of abuse was performed in each urine specimen and the following criteria are applied for positive drug detection.

| Panel of Immunoassay | Cut-off for positive | Lowest detection limit |
|------------------------|------------------------|------------------------|
| (Abbott AxSym® System) | drug detection | |
| Amphetamines | <u>></u> 1000 ng/mL | 100.00 ng/mL |
| Barbiturates | <u>></u> 200 ng/mL | 60.00 ng/mL |
| Benzodiazepines | <u>></u> 200 ng/mL | 40.00 ng/mL |
| Cannabinoids | <u>></u> 50 ng/mL | 13.00 ng/mL |
| Cocaine | <u>></u> 300 ng/mL | 30.00 ng/mL |
| Methadone | <u>></u> 250 ng/mL | 100.00 ng/mL |
| Opiates | <u>></u> 300 ng/mL | 50.00 ng/mL |
| | | |

*Signals above the lowest detection limit but below the cut-off value for the corresponding assay are regarded as negative results.

| Drug category | Drug item detected | Drug category | Drug item detected | |
|-----------------|---------------------------------|-------------------|--------------------|-----------------|
| Amphetamines | Methamphetamine /Amphetamine | Therapeutic drugs | Amisulpride | Ofloxacin |
| Analgesics | Tramadol | | Amitriptyline | Olanzapine |
| Barbiturates | Barbiturates | | Atenolol | Perindopril |
| Benzodiazepines | Diazepam | | Atropine | Propranolol |
| | Flunitrazepam | | Benzhexol | Quetiapine |
| | Midazolam | | Carbamazepine | Ranitidine |
| | Nitrazepam | | Chlorpromazine | Risperidone |
| Cannabis | Cannabinoids | | Cimetidine | Sertraline |
| Cocaine | Cocaine | | Citalopram | Sildenafil |
| Cough medicine | Brompheniramine | | Clarithromycin | Sulpiride |
| | Cetirizine | | Clozapine | Trazodone |
| | Chlorpheniramine | | Desipramine | Trifluoperazine |
| | Codeine | | Diphenhydramine | Trimipramine |
| | Dextromethorphan | | Dothiepin | Venlafaxine |
| | Dihydrocodeine | | Doxepin | Zuclopenthixol |
| | Hydrocodone | | Etoricoxib | |
| | Hydromorphone | | Famotidine | |
| | Methylephedrine | | Fluconazole | |
| | Promethazine | | Fluoxetine | |
| | Propoxyphene | | Glibenclamide | |
| | Pseudoephedrine/Ephedrine | | Gliclazide | |
| Hypnotics | Zolpidem | | Haloperidol | |
| | Zopiclone | | Lidocaine | |
| Opiates | Heroin | | Metoclopramide | |
| Ketamine | Ketamine | | Metoprolol | |
| MDMA | HMMA | | Metronidazole | |
| | MDA | | Mianserin | |
| | MDMA | | Mirtazapine | |
| Methadone | Methadone | | Nortriptyline | |
| | Subtotal: 29 | | Subtotal: 44 | |

2.2.1.2 Summary of drug items detected in all urine samples by UPLC-TOF-MS

*A total of 73 drug items (including both drugs of abuse or therapeutic agents) were identified from 2,783 parent drugs and metabolites detected by UPLC-TOF-MS in 454 urine samples.

2.2.1.3 Summary of number of drug of abuse items detected in urine

| Number of groups of | Femal | e | Male | | Total | |
|---------------------|-------|---------|------|---------|-------|---------|
| drug of abuse items | | | | | | |
| | Ν | % | Ν | % | Ν | % |
| 0 | 59 | (33%) | 79 | (28.7%) | 138 | (30.4%) |
| 1 | 74 | (41.3%) | 122 | (44.4%) | 196 | (43.2%) |
| 2 | 40 | (22.3%) | 52 | (18.9%) | 92 | (20.3%) |
| 3 | 5 | (2.8%) | 16 | (5.8%) | 21 | (4.6%) |
| 4 | 1 | (0.6%) | 6 | (2.2%) | 7 | (1.5%) |
| Grand Total | 179 | (100%) | 275 | (100%) | 454 | (100%) |

Number of groups of drug of abuse items detected in urine (by sex)

Number of groups of drug of abuse items detected in urine (by age)

| Number of groups | | Age group (Years) | | | | | | |
|------------------|-------|-------------------|-------|-------|-------|-------|-----|---------|
| of drug of abuse | 10-19 | 20-29 | 30-39 | 40-49 | 50-59 | 60-70 | N | % |
| items | | | | | | | | |
| 0 | 30 | 39 | 40 | 20 | 9 | | 138 | (30.4%) |
| 1 | 14 | 56 | 65 | 35 | 23 | 3 | 196 | (43.2%) |
| 2 | 3 | 21 | 39 | 24 | 4 | 1 | 92 | (20.3%) |
| 3 | 2 | 2 | 9 | 3 | 5 | | 21 | (4.6%) |
| 4 | 1 | | 6 | | | | 7 | (1.5%) |
| Grand Total | 50 | 118 | 159 | 82 | 41 | 4 | 454 | (100%) |

*Note: Drug items under the category of therapeutic drugs and methadone are not included in this table.

*Number of group of drug item detected in individual urine sample ranged from 0 to 4.

*Proportion of clean urine (i.e. no drug of abuse item detected) within an individual age-group decreased among increasing age. [60% in 10-19 years, 33.1% in 20-21 years, 25.2% in 30-39 years, 24.4% in 40-49 years, 22% in 50-59 years and 0% in 60-70 years.]

2.2.1.4 Patterns of group of drug of abuse item detected

| Group of drug item | Female | | Male | Male | | |
|--------------------|--------|---------|-------|---------|--------|---------|
| | (N=1) | 79) | (N=2) | 75) | (N=454 | .) |
| | | % | | % | | % |
| Cough medicine | 37 | (20.7%) | 84 | (30.5%) | 121 | (26.7%) |
| Hypnotics | 43 | (24%) | 40 | (14.5%) | 83 | (18.3%) |
| Ketamine | 30 | (16.8%) | 44 | (16%) | 74 | (16.3%) |
| Opiates | 25 | (14%) | 46 | (16.7%) | 71 | (15.6%) |
| Benzo | 22 | (12.3%) | 39 | (14.2%) | 61 | (13.4%) |
| Amphe | 10 | (5.6%) | 33 | (12%) | 43 | (9.5%) |
| Cocaine | 4 | (2.2%) | 1 | (0.4%) | 5 | (1.1%) |
| Cannabis | | | 4 | (1.5%) | 4 | (0.9%) |
| MDMA | 1 | (0.6%) | 1 | (0.4%) | 2 | (0.4%) |
| Barbiturates | | | 1 | (0.4%) | 1 | (0.2%) |
| Analgesic | | | 1 | (0.4%) | 1 | (0.2%) |
| Grand Total | 172 | | 294 | | 466 | |

Summary of group of drug of abuse item detected in urine (by sex)
 (Data shown in descending order of the frequency detected)

*Note: Methadone was not counted as drug of abuse category in this table.

*Percentage is calculated based on the total number (N) of subjects of the corresponding sex and the total number of recruited subjects.

*Cough medicine was the most frequent group of drug of abuse detected in the urine of our study population. Hypnotics was the second most frequent group of drug of abuse detected, followed by ketamine, opiates and benzodiazepines.

*Higher frequency of hypnotics was detected in urine of female subjects, but higher frequency of cough medicine was detected in male subjects.

| | Age group (Years) | | | | | | | | | | | | Grand Total (N=454) | |
|-------------------------|-------------------|-------|-------|---------|-------|---------|------|---------|------|---------|-----|-------|------------------------|---------|
| Groups of drug items | 10-1 | .9 | 20-29 |) | 30-39 |) | 40-4 | 19 | 50-5 | 59 | 60- | 70 | | |
| items | (N= | 50) | (N=1) | 18) | (N=1 | 59) | (N= | 32) | (N=4 | 41) | (N= | =4) | | |
| | Ν | % | N | % | Ν | % | Ν | % | Ν | % | Ν | % | N | % |
| Cough medicine | 6 | (12%) | 24 | (20.3%) | 64 | (40.3%) | 20 | (24.4%) | 6 | (14.6%) | 1 | (25%) | 121 | (26.7%) |
| Hypnotics | 2 | (4%) | 11 | (9.3%) | 31 | (19.5%) | 23 | (28%) | 13 | (31.7%) | 3 | (75%) | 83 | (18.3%) |
| Ketamine | 11 | (22%) | 48 | (40.7%) | 15 | (9.4%) | | | | | | | 74 | (16.3%) |
| Opiates | | | 3 | (2.5%) | 27 | (17%) | 23 | (28%) | 17 | (41.5%) | 1 | (25%) | 71 | (15.6%) |
| Benzo | 1 | (2%) | 7 | (5.9%) | 27 | (17%) | 17 | (20.7%) | 8 | (19.5%) | 1 | (25%) | 61 | (13.4%) |
| Amphe | 4 | (8%) | 9 | (7.6%) | 20 | (12.6%) | 9 | (11%) | 1 | (2.4%) | | | 43 | (9.5%) |
| Cocaine | 2 | (4%) | 2 | (1.7%) | 1 | (0.6%) | | | | | | | 5 | (1.1%) |
| Cannabis | | | | | 3 | (1.9%) | 1 | (1.2%) | | | | | 4 | (0.9%) |
| MDMA | 2 | (4%) | | | | | | | | | | | 2 | (0.4%) |
| Barbiturates | | | | | | | | | 1 | (2.4%) | | | 1 | (0.2%) |
| Analgesic | | | | | 1 | (0.6%) | | | | | | | 1 | (0.2%) |
| Grand Total | 28 | | 104 | | 189 | | 93 | | 46 | | 6 | | 466 | |

Summary of group of drug of abuse item detected in urine (by age) (Data shown in descending order of the frequency detected)

*Note: One or more groups of drug items were detected in individual urine sample. Figures are presented as the frequency of a particular drug item detected in corresponding age group. Percentage is calculated based on the total number (N) of subjects of the corresponding age group and the total number of recruited subjects.

*Patterns of drugs detected were different in different age groups. Cough medicine and ketamine were detected in the younger age groups more frequently while opiates and hypnotics were detected in the older age groups more frequently.

| Ingredients of cough medicine detected in urine | Grand To | tal |
|-------------------------------------------------|----------|-------|
| | (N=121) | |
| | | % |
| Pseudoephedrine/Ephedrine | 89 | 73.6% |
| Promethazine | 88 | 72.7% |
| Codeine | 67 | 55.4% |
| Hydrocodone | 48 | 39.7% |
| Chlorpheniramine | 46 | 38% |
| Dextromethorphan | 24 | 19.8% |
| Propoxyphene | 23 | 19% |
| Brompheniramine | 16 | 13.2% |
| Dihydrocodeine | 5 | 4.1% |
| Diphenhydramine | 3 | 2.5% |
| Methylephedrine | 2 | 1.7% |
| Cetirizine | 1 | 0.8% |
| Hydromorphone | 1 | 0.8% |
| Grand Total | 413 | |

Common ingredients of cough medicine detected
 (Data shown in descending order of the frequency detected)

*Note: Urine samples with detection of two or more cough medicine ingredients were counted as positive detection for cough medicine except the detection of pseudoephedrine/ephedrine or codeine alone would also be counted as positive.

- *Note: One or more ingredients of cough medicine could be detected in the same urine sample. Percentage is calculated based on the total number (N) of urine samples with positive detection of cough medicine.
- *Note: Promethazine is a first-generation H1 receptor antagonist of the phenothiazine chemical class. It has a strong sedative effect, as well as anti-motion-sickness, anti-emetic, and anti-cholinergic effects. Fifteen urine samples with detection of promethazine with or without other therapeutic drugs were not counted in the above table as it can be a prescribed hypnotic in SAC for therapeutic use.
- *A total of 13 drug items in the category of cough medicine were detected in our study population
- *Pseudoephedrine/ephedrine, promethazine, codeine, hydrocodone and chlorpheniramine are the five most common ingredients of cough medicine detected.

2.2.1.5 Unexpected substances detected in urine samples

- Novel drugs or impurities
- *No unidentified peak of unknown substances was detected in the 454 urine samples so far.
- *High prevalence of cough medicine was detected in urine of our study subjects. Ingredients of cough medicine were detected in 121 urine samples (26.7% of 454 subjects).
- *However, a total of 376 (82.8%) of 454 subjects did not report cough medicine in their last use, of which, 64 subjects (17% of 376 subjects) turned out to have positive findings of ingredients of cough medicine in their urine.
- *Of these 64 subjects, the six most frequently reported drug items in their last drug use were hypnotics (24 subject, 37.5%), ketamine (14 subjects, 21.9%), opiates (14 subjects, 21.9%), amphetamines (8 subjects, 12.5%) and benzodiazepines (5 subjects, 7.8%). [Note: Percentage was calculated based on number of subjects who have negative history but positive detection of cough medicine, i.e. N=64. Subjects could report more than one drug item during their last use.]

2.2.2 Comparison of conventional and new techniques

2.2.2.1 Scoring system

- Compare the efficiency between conventional (HPLC-UV/ immunoassays) and new (UPLC-TOF/MS) techniques
 - Other than drugs caffeine, nicotine and cotinine, count the number of drugs identified in each urine sample by both conventional (C) and new (N) techniques.
 - 2. Number of drugs identified by conventional techniques (C) is defined as the number of drugs identified by HPLC-UV and number of additional drugs identified by immunoassay.
 - 3. Determine the ratio of C/N of each urine sample.
 - 4. The presence of drugs in both libraries should not be considered.
 - 5. Calculate the mean C/N ratio. If the mean ratio is <1, the new technique identifies more drugs than conventional technique. If the mean ratio is >1, the conventional technique identifies more drugs than the new technique.

- > Performance of the New for replacement of Conventional technique
 - 1. Compare the identified drugs between the New and Conventional techniques.
 - 2. For drugs identified by the Conventional technique but not the New, add +1 point to Deficiency score (D) for each drug.
 - 3. For drugs identified by the New technique but not the Conventional, add +1 point to Improvement score (I) for each drug.
 - 4. Calculate the D/I ratio. If the D/I ratio is <1, the New technique has better overall performance than the Conventional technique in terms of identification of drug item by detection of either parent drug or metabolites. If the D/I ratio is >1, the Conventional technique has better performance and the New technique has missed drugs that are identified by the Conventional technique.

2.2.2.2 Results of the comparison data

| Summary of performance of new and conventional techniques | |
|----------------------------------------------------------------------------|------|
| Number of urine sample analyzed by both techniques | 67 |
| Total number of drug items identified by new technique (sum of N) | 171 |
| Total number of drug items identified by conventional technique (sum of C) | 147 |
| Mean C/N ratio | 0.77 |
| Deficiency score (D) | 22 |
| Improvement score (I) | 46 |
| D/I ratio | 0.48 |
| | |

2.2.2.3 Summary of the comparison data

Sixty-seven urine samples were analyzed by both of the new (UPLC-TOF-MS) and conventional (HPLC-UV + Immunoassay) techniques. The total number of drug items identified by the new technique is greater than that of the conventional technique. Both the mean C/N ratio and the D/I ratio are less than one. From our current data, the new technique was able to enhance the drug detection when compared with the conventional method. Therefore, the urine analysis results generated by UPLC-TOF-MS provide us useful and reliable information for the investigation of the pattern of substance abuse among our study population.

2.3 Comparison of survey and urine analysis results

2.3.1 Patterns of substance abuse

Five most common drug of abuse items reported (last drug use) or detected in our study population (N = 454)

| Results of survey | | | Results of urine ana | Results of urine analysis | | | |
|-------------------|-----|---------|----------------------|---------------------------|---------|--|--|
| | Ν | % | | Ν | % | | |
| Ketamine | 113 | (24.9%) | Cough medicine | 121 | (26.7%) | | |
| Hypnotics | 83 | (18.3%) | Hypnotics | 83 | (18.3%) | | |
| Opiates | 82 | (18.1%) | Ketamine | 74 | (16.3%) | | |
| Cough medicine | 78 | (17.2%) | Opiates | 71 | (15.6%) | | |
| Amphetamines | 72 | (15.9%) | Benzodiazepines | 61 | (13.4%) | | |
| | | | | | | | |

Five most common drug of abuse items in female (N = 179)

| Results of survey | Results of urine analysis | | | | |
|-------------------|---------------------------|---------|-----------------|----|---------|
| | Ν | % | | Ν | % |
| Hypnotics | 48 | (26.8%) | Hypnotics | 43 | (24%) |
| Ketamine | 44 | (24.6%) | Cough medicine | 37 | (20.7%) |
| Amphetamines | 43 | (24%) | Ketamine | 30 | (16.8%) |
| Opiates | 29 | (16.2%) | Opiates | 25 | (14.0%) |
| Benzodiazepines | 15 | (8.4%) | Benzodiazepines | 22 | (12.3%) |
| | | | | | |

Five most common drug of abuse items in male (N = 275)

| Results of survey | | | Results of urine analy | /sis | |
|-------------------|----|---------|------------------------|------|---------|
| | Ν | % | | Ν | % |
| Ketamine | 69 | (25.1%) | Cough medicine | 84 | (30.5%) |
| Cough medicine | 64 | (23.3%) | Opiates | 46 | (16.7%) |
| Opiates | 53 | (19.3%) | Ketamine | 44 | (16.0%) |
| Hypnotics | 35 | (12.7%) | Hypnotics | 40 | (14.5%) |
| Amphetamines | 29 | (10.5%) | Benzodiazepines | 39 | (14.2%) |
| | | | | | |

- Pattern of drug of abuse items reported (last drug use) and detected in urine of subjects by different age group (Data shown in descending order of the frequency detected)
- *Note: One or more groups of drug items were detected in individual urine sample. Figures are presented as the frequency of a particular drug item detected and percentage is calculated based on the number (N) of subjects in a particular age group.

*Different prevalence of drug use is shown in the following tables.

(1) Age of 10 – 19 years (N = 50)

| Results of survey | | | Results of urine analysis | | | |
|-------------------|----|-------|---------------------------|----|-------|--|
| | Ν | % | | Ν | % | |
| Ketamine | 27 | (54%) | Ketamine | 11 | (22%) | |
| Amphetamines | 19 | (38%) | Cough medicine | 6 | (12%) | |
| Cocaine | 5 | (10%) | Amphetamines | 4 | (8%) | |
| Benzodiazepines | 2 | (4%) | Hypnotics | 2 | (4%) | |
| Mandrax | 2 | (4%) | Cocaine | 2 | (4%) | |
| Not specified | 2 | (4%) | MDMA | 2 | (4%) | |
| | | | | | | |

(2) Age of 20 – 29 years (N = 118)

| Results of survey | | | Results of urine analysis | | | |
|-------------------|----|---------|---------------------------|----|---------|--|
| | Ν | % | | Ν | % | |
| Ketamine | 69 | (58.5%) | Ketamine | 48 | (40.7%) | |
| Amphetamines | 23 | (19.5%) | Cough medicine | 24 | (20.3%) | |
| Cough medicine | 13 | (11%) | Hypnotics | 11 | (9.3%) | |
| Hypnotics | 8 | (6.8%) | Amphetamines | 9 | (7.6%) | |
| Opiates | 6 | (5.1%) | Benzodiazepines | 7 | (5.9%) | |
| | | | | | | |

(3) Age of 30 – 39 years (N = 159)

| Results of survey | | | Results of urine analysis | | | |
|-------------------|----|---------|---------------------------|----|---------|--|
| | N | % | | Ν | % | |
| Cough medicine | 52 | (32.7%) | Cough medicine | 64 | (40.3%) | |
| Opiates | 36 | (22.6%) | Hypnotics | 31 | (19.5%) | |
| Amphetamines | 22 | (13.8%) | Opiates | 27 | (17%) | |
| Hypnotics | 18 | (11.3%) | Benzodiazepines | 27 | (17%) | |
| Ketamine | 16 | (10.1%) | Amphetamines | 20 | (12.6%) | |
| | | | | | | |

(4) Age of 40 – 49 years (N = 82)

| Results of survey | | | Results of urine analysis | | | |
|-------------------|----|---------|---------------------------|----|---------|--|
| | Ν | % | | Ν | % | |
| Hypnotics | 33 | (40.2%) | Hypnotics | 23 | (28%) | |
| Opiates | 24 | (29.3%) | Opiates | 23 | (28%) | |
| Cough medicine | 11 | (13.4%) | Cough medicine | 20 | (24.4%) | |
| Cannabis | 9 | (11%) | Benzodiazepines | 17 | (20.7%) | |
| Amphetamines | 8 | (9.8%) | Amphetamines | | (11%) | |
| | | | | | | |

(5) Age of 50 – 59 years (N = 41)

| Results of survey | | | Results of urine and | Results of urine analysis | | | | |
|-------------------|----|---------|----------------------|---------------------------|---------|--|--|--|
| | N | % | | Ν | % | | | |
| Hypnotics | 21 | (51.2%) | Opiates | 17 | (41.5%) | | | |
| Opiates | 14 | (34.1%) | Hypnotics | 13 | (31.7%) | | | |
| Benzodiazepines | 4 | (9.8%) | Benzodiazepines | 8 | (19.5%) | | | |
| Cannabis | 2 | (4.9%) | Cough medicine | 6 | (14.6%) | | | |
| Cough medicine | 1 | (2.4%) | Amphetamines | 1 | (2.4%) | | | |
| | | | Barbiturates | 1 | (2.4%) | | | |
| | | | | | | | | |

(6) Age of 60 – 70 years (N = 4)

| Results of survey | | | Results of urine analysis | | | |
|-------------------|---|-------|---------------------------|---|-------|--|
| | N | % | | Ν | % | |
| Hypnotics | 2 | (50%) | Hypnotics | 3 | (75%) | |
| Opiates | 2 | (50%) | Cough medicine | 1 | (25%) | |
| | | | Opiates | 1 | (25%) | |
| | | | Benzodiazepines | 1 | (25%) | |
| | | | | | | |

Pattern of drug of abuse items detected in study subjects recruited from different regions of the new territories

(Data shown in descending order of the frequency detected of the grand total.)

*Note: One or more groups of drug items were detected in individual urine sample. Figures are presented as the frequency of a particular drug item detected and percentages are calculated based on the number (N) of subjects of the corresponding sex and corresponding cluster.

| Drug items detected in urine | Female (N=73) | | | Male (N=127) | | l 200) |
|------------------------------|------------------|---------|-----|-----------------|-----|-----------|
| | | % | | % | | % |
| Cough medicine | 11 | (15.1%) | 40 | (31.5%) | 51 | (25.5%) |
| Ketamine | 16 | (21.9%) | 23 | (18.1%) | 39 | (19.5%) |
| Hypnotics | 16 | (21.9%) | 22 | (17.3%) | 38 | (19%) |
| Benzodiazepines | 10 | (13.7%) | 24 | (18.9%) | 34 | (17%) |
| Opiates | 8 | (11%) | 16 | (12.6%) | 24 | (12%) |
| Amphetamines | 6 | (8.2%) | 15 | (11.8%) | 21 | (10.5%) |
| Cocaine | 4 | (5.5%) | 1 | (0.8%) | 5 | (2.5%) |
| MDMA | 1 | (1.4%) | 1 | (0.8%) | 2 | (1%) |
| Grand Total | 72 | | 142 | | 214 | |

(1) Subjects recruited from NTEC

(2) Subjects recruited from NTWC

| Duug itoma datastad in uning | Female | | Male | | Total | |
|------------------------------|--------|---------|------|---------|-------|---------|
| Drug items detected in urine | (N=1 | .06) | (N=1 | (N=148) | | .54) |
| | | % | | % | | % |
| Cough medicine | 26 | (24.5%) | 43 | (29.1%) | 69 | (27.2%) |
| Hypnotics | 29 | (27.4%) | 18 | (12.2%) | 47 | (18.5%) |
| Opiates | 17 | (16%) | 29 | (19.6%) | 46 | (18.1%) |
| Ketamine | 15 | (14.2%) | 19 | (12.8%) | 34 | (13.4%) |
| Benzodiazepines | 12 | (11.3%) | 14 | (9.5%) | 26 | (10.2%) |
| Amphetamines | 5 | (4.7%) | 18 | (12.2%) | 23 | (9.1%) |
| Cannabis | | | 4 | (2.7%) | 4 | (1.6%) |
| MDMA | | | 1 | (0.7%) | 1 | (0.4%) |
| Barbiturates | | | 1 | (0.7%) | 1 | (0.4%) |
| Analgesic | | | 1 | (0.7%) | 1 | (0.4%) |
| Grand Total | 104 | | 148 | | 252 | |

(3) Comparing the five most common drug of abuse items detected in subjects recruited from NTEC and NTWC

| NTEC | (N=: | 200) | NTWC | (N= | (N=254) | |
|-----------------|------|---------|-----------------|-----|---------|--|
| | Ν | % | | Ν | % | |
| Cough medicine | 51 | (25.5%) | Cough medicine | 69 | (27.2%) | |
| Ketamine | 39 | (19.5%) | Hypnotics | 47 | (18.5%) | |
| Hypnotics | 38 | (19%) | Opiates | 46 | (18.1%) | |
| Benzodiazepines | 34 | (17%) | Ketamine | 34 | (13.4%) | |
| Opiates | 24 | (12%) | Benzodiazepines | 26 | (10.2%) | |
| | | | | | | |

*The five most common items detected in urine were same but with different prevalence in the two subgroups of study subjects recruited from different regions of the new territories.

2.3.2 Comparison of urine analysis results with history of last drug use

| Time of last drug | No. | of dru | g iten | n last u | sed | No. | of grou | up of d | rug det | ected | Total | |
|------------------------|------|--------|--------|----------|-----|-------|---------|---------|---------|-------|-------|---------|
| use | repo | orted | | | | in ur | ine | | | | (N=4 | 54) |
| | NS | 1 | 2 | 3 | 4 | 0 | 1 | 2 | 3 | 4 | | |
| < 1 Day | | 88 | 7 | | 1 | 3 | 51 | 30 | 9 | 3 | 96 | (21.1%) |
| 1 day to <= 1 week | 1 | 116 | 11 | 1 | 2 | 9 | 74 | 37 | 8 | 3 | 131 | (28.9%) |
| > 1 week to <= 1 month | 1 | 54 | 3 | | | 23 | 25 | 6 | 3 | 1 | 58 | (12.8%) |
| > 1 month to <= 1 year | | 93 | 12 | 2 | 1 | 69 | 23 | 15 | 1 | | 108 | (23.8%) |
| > 1 year | 1 | 42 | | 1 | | 28 | 14 | 2 | | | 44 | (9.7%) |
| Not specified | 10 | 4 | 1 | | | 5 | 8 | 2 | | | 15 | (3.3%) |
| Quitted | 2 | | | | | 1 | 1 | | | | 2 | (0.4%) |
| Grand Total | 15 | 397 | 34 | 4 | 4 | 138 | 196 | 92 | 21 | 7 | 454 | (100%) |

Number of drug use reported and number of drug detected in urine with reference to the timing of last drug use

*Note: NS = Not specified from the history

*From the survey result, most subjects reported using one drug only during their last use.

- *Out of 454 subjects, 227 subjects reported to have last drug use within 1 week's time. In this group, when comparing the history to the urine results in general, 204 (89.9%) of 227 subjects reported single drug use only and 22 (9.7%) of 227 subjects reported multiple drug use while 125 (55.1%) of these 227 urine samples detected one drug of abuse item and 90 (39.6%) of these 227 urine samples detected two or more drug of abuse items. This observation suggested an underreporting of the number of drug abused which may be intentional (i.e. the subject deliberately hide the genuine drug-using habit) or incidental (i.e. the subject did not know the impurities in the substances he/she took).
- *Out of 454 subjects, 210 subjects reported to have last drug use greater than one week's time. Because of the limitation of the urine analysis for sample taken after one week of the last use, no drug of abuse item should be detected in these urine samples and any presence of drug of abuse items in urine could signify recent drug use less than 1 week. In this group of subjects, 120 (57.1%) of 210 subjects have negative urine result while 90 (42.9%) of 210 subjects have positive urine results. Twenty-eight (31.1%) of the 90 subjects who have positive urine results even have two or more drug of abuse items detected in their urine samples.

- * Forty-four subjects reported no drug use for one or more years (up to 10 years), 16 (36.4%) of which still have positive urine results. Cough medicine (5 samples), hypnotics (4 samples), opiates (4 samples), benzodiazepines (4 samples) and amphetamines (1 sample) were detected in this 16 urine samples.
- Concordance of individual's history of last use and corresponding urine result

Six commonly reported/detected drugs are analyzed.

| History of last use | Urine detection by UPLC-TOF-MS | | | |
|---------------------|--------------------------------|----------|-------------|--|
| History of last use | Positive | Negative | Grand Total | |
| Positive | А | В | A + B | |
| Negative | С | D | C + D | |
| Grand Total | A + C | B + D | 454 | |

The results are tabulated as below.

*Note:

- A = Number of subjects who reported to have positive drug use and had positive urine result
- B = Number of subjects who reported to have positive drug use but had negative urine result
- C = Number of subjects who reported no drug use but had positive urine result
- D = Number of subjects who reported no drug use and had negative urine result
- A + B = Number of subjects who had positive drug use history
- A + C = Number of subjects who had positive urine drug detection
- *A and D signify that the history and urine result are concordant.
- *B and C signify that the history and urine result are discordant.

(1) Ketamine

All subjects

| History of last use | Urine detection l | | |
|---------------------|-------------------|----------|-------------|
| HISTORY OF IAST USE | Positive | Negative | Grand Total |
| Positive | 66 | 47 | 113 |
| Negative | 8 | 333 | 341 |
| Grand Total | 74 | 380 | 454 |

*A + D = 399 (Concordance rate = 87.9%)

Subjects reported last drug use of < one week</p>

| liston, of last use | Urine detection by UPLC-TOF-MS | | | | |
|---------------------|--------------------------------|----------|-------------|--|--|
| History of last use | Positive | Negative | Grand Total | | |
| Positive | 55 | 3 | 58 | | |
| Negative | 6 | 163 | 169 | | |
| Grand Total | 61 | 166 | 227 | | |

*A + D = 218 (Concordance rate = 96%)

*If one week is taken as the window period of ketamine detection in urine, the positive detection rate by UPLC-TOF-MS in subjects with positive history of ketamine use is 94.8% (55 of 58 subjects). Of the 113 subjects who reported ketamine in their last use, 58 subjects reported their last use within one week's time. Fifty-five of the 58 urine samples detected ketamine. Of the three urine samples negative for ketamine, cough medicine instead of ketamine was detected in one urine sample while the other two urine samples did not detect any drug.

*Forty-seven of 113 subjects who reported ketamine in their last use have negative urine finding for ketamine. Forty-three of 47 subjects reported their use of greater than one week's time, 1 reported less than one day, 2 reported one week or less and 1 subject did not specified the time of his/her last use. Among these 43 subjects, 28 subjects have no drug detected in their urine samples, 2 subjects have therapeutic drugs detected and 1 subject has methadone detected, Six of the 43 subjects have one drug of abuse item detected and cough medicine was detected in five of these six urine samples.

- *Seventy-four (16.3%) of 454 urine samples have positive findings of ketamine. Ten (13.5%) of the 74 urine samples also had positive findings of cough medicine.
- *Eight (1.8%) of 454 subjects who were under-reporting their ketamine abuse in their last use were discovered by the positive urine findings.

(2) Amphetamines

➢ All subjects

| History of last use | Urine detection l | | |
|---------------------|-------------------|----------|-------------|
| HISTORY OF IAST USE | Positive | Negative | Grand Total |
| Positive | 22 | 50 | 72 |
| Negative | 21 | 361 | 382 |
| Grand Total | 43 | 411 | 454 |

*A + D = 383 (Concordance rate = 84.4%)

Subjects reported last drug use of < one week

| History of last use | Urine detection l | | |
|---------------------|-------------------|----------|-------------|
| History of last use | Positive | Negative | Grand Total |
| Positive | 14 | 5 | 19 |
| Negative | 16 | 192 | 208 |
| Grand Total | 30 | 197 | 227 |

*A + D = 206 (Concordance rate = 90.7%)

*If one week is taken as the window period of amphetamines detection in urine, the positive detection rate by UPLC-TOF-MS in subjects with positive history of amphetamines use is 73.7%% (14 of 19 subjects). Of the 72 subjects who reported amphetamines in their last use, 19 subjects reported their last use within one week's time. Fourteen of the 19 urine samples detected amphetamines and all of these 14 urine samples were collected within 4 days of the reported last use. Of the 5 urine samples negative for amphetamines, 3 were collected 7 days after the reported last use and 1 was collected 4 days after. For the one negative urine sample that was collected 1 day after the reported last use, benzodiazepine and cough medicine instead of amphetamines were detected in this urine sample. If the urine samples are taken within 4 days of last use, the positive drug detection rate by UPLC-TOF-MS is 87.5% (14 of 16 subjects). The detection rate would be

increased to 92.3% (12 of 13 subjects) if 3 days is taken as a cut-off time for the urine sampling.

- *Fifty of 72 subjects who reported amphetamines in their last use have negative urine finding for amphetamines. Forty-five of 50 subjects reported their use of greater than one week's time, of which, 11 subjects reported greater than one week but less than one month, 26 subjects reported greater than one month but less than one year, 6 subjects reported greater than one year's time and 2 subjects did not specified the time of their last use. Thirty-five of 45 subjects have no drug of abuse items detected in their urine sample. Ten (22.2%) of 45 subjects have one or more drug of abuse items detected and cough medicine and hypnotics were detected in 4 of these 10 urine samples. Other drugs of abuse detected include opiates (in 2 urine samples) and benzodiazepines (in 2 urine samples).
- * Forty-three (9.5%) of 454 urine samples have positive findings of amphetamines. Overall, 21 subjects (4.6% of total 454 subjects; 48.8% of 43 subjects with positive urine findings) who were under-reporting their amphetamines abuse in their last use were discovered by the positive urine findings.

(3) Hypnotics

All subjects

| History of last use | Urine detection l | | |
|---------------------|-------------------|----------|-------------|
| History of last use | Positive | Negative | Grand Total |
| Positive | 47 | 36 | 83 |
| Negative | 36 | 335 | 371 |
| Grand Total | 83 | 371 | 454 |

*A + D = 382 (Concordance rate = 84.1%)

Subjects reported last drug use of < one week</p>

| History of last use | Urine detection l | | |
|---------------------|-------------------|----------|-------------|
| HISTORY OF Idst use | Positive | Negative | Grand Total |
| Positive | 43 | 17 | 60 |
| Negative | 23 | 144 | 167 |
| Grand Total | 66 | 161 | 227 |

*A + D = 187 (Concordance rate = 82.4%)

- *If one week is taken as the window period of hypnotics (e.g. zopiclone and zolpidem) detection in urine, the positive detection rate by UPLC-TOF-MS in subjects with positive history of hypnotics use is 71.7% (43 of 60 subjects). Of the 83 subjects who reported hypnotics in their last use, 60 subjects reported their last use within one week's time. Forty subjects of the 43 subjects with positive urine results reported their last use of one day or less, 2 subjects reported two days and 1 subject reported five days.
- *Thirty-six of 83 subjects who reported hypnotics in their last use have negative urine finding for hypnotics. Eighteen of the 36 subjects reported their last use of greater than one week's time and one subject did not specified the time of last drug use. Fourteen (77.8%) of the 18 subjects had positive urine results for other drugs, such as benzodiazepines (in 9 samples), cough medicine (in 4 samples) and opiates (in 1 sample).
- *Eighty-three of 454 urine samples from 43 females and 40 males have positive findings of hypnotics. Seventy-three urine samples detected zopiclone while 10 urine samples detected zolpidem. Majority, 31 (37.3%) and 23 (27.7%) of the 83 subjects, were in the age groups of 30-39 years and 40-49 years respectively. Sixty-three (75.9%) of the 83 subjects reported either physical or psychiatric complications from hypnotics use. Most commonly complained psychiatry complications include impairment of memory (in 41 subjects), depression (in 29 subjects), psychosis (in 16 subjects) and hallucination (in 15 subjects). Four subjects with chronic hypnotics use reported history of suicidal attempt. Physical complications reported were non-specific or related to withdrawal symptoms of chronic hypnotics use.
- *Thirty-six subjects (7.9% of total 454 subjects; 43.4% of 83 subjects with positive urine findings; 11 females and 25 males) who were under-reporting their hypnotics abuse in their last use were discovered by the positive urine findings. Of the 36 subjects, 19 (52.7%) subjects were in the age group of 30-39 years.

(4) Cough medicine

All subjects

| History of last use | Urine detection I | | |
|---------------------|-------------------|----------|-------------|
| History of last use | Positive | Negative | Grand Total |
| Positive | 57 | 21 | 78 |
| Negative | 64 | 312 | 376 |
| Grand Total | 121 | 333 | 454 |

*A + D = 369 (Concordance rate = 81.3%)

Subjects reported last drug use of < one week

| listen of last up | Urine detection by UPLC-TOF-MS | | | |
|---------------------|--------------------------------|----------|-------------|--|
| History of last use | Positive | Negative | Grand Total | |
| Positive | 45 | 4 | 49 | |
| Negative | 35 | 143 | 178 | |
| Grand Total | 80 | 147 | 227 | |

*A + D = 188 (Concordance rate = 82.8%)

- *Twenty-one of 78 subjects who reported cough medicine in their last use have negative urine finding for cough medicine. Four subjects reported their last use of greater than one year, 10 subjects reported greater than one month, 3 subjects reported greater than one week and 4 subjects reported less than one week. Hypnotics and benzodiazepines were detected in 2 of the 4 subjects who reported last use of cough medicine within one week but have negative detection for cough medicine in urine.
- *One hundred and twenty-one of 454 urine samples have positive findings of cough medicine. High unreported rate, 64 (14.1%) of total (52.9% of 121 subjects with positive urine findings) was discovered by the positive urine findings. Of the 64 subjects, 35 (54.7%) reported their last use of drugs within one week's time. Of the 35 urine samples, 30 (85.7%) detected multiple drug of abuse items despite cough medicine. Seventeen and 18 subjects were female and male, respectively. Of the 35 subjects, 22

^{*}If one week is taken as the window period of cough medicine detection in urine, the positive detection rate by UPLC-TOF-MS in subjects with positive history of ketamine use is 91.8% (45 of 49 subjects). Of the 78 subjects who reported cough medicine in their last use, 49 subjects reported their last use within one week's time.

(62.9%) were in the young age groups of less than 40 years (12 subjects in 30-39 years; 8 subjects in 20-29 years; 2 subjects in 10-19 years).

(5) Opiates (Heroin)

All subjects

| History of last use | Urine detection l | | |
|---------------------|-------------------|----------|-------------|
| History of last use | Positive | Negative | Grand Total |
| Positive | 52 | 30 | 82 |
| Negative | 19 | 353 | 372 |
| Grand Total | 71 | 383 | 454 |

*A + D = 405 (Concordance rate = 89.2%)

Subjects reported last drug use of < one week

| History of last use | Urine detection by UPLC-TOF-MS | | |
|---------------------|--------------------------------|----------|-------------|
| | Positive | Negative | Grand Total |
| Positive | 41 | 2 | 43 |
| Negative | 11 | 173 | 184 |
| Grand Total | 52 | 175 | 227 |

*A + D = 214 (Concordance rate = 94.3%)

*If one week is taken as the window period of opiates [Heroin] detection in urine, the positive detection rate by UPLC-TOF-MS in subjects with positive history of heroin use is 95.3% (41 of 43 subjects). Of the 82 subjects who reported heroin in their last use, 43 subjects reported their last use within one week's time. Of the 43 urine samples, 41 detected Heroin and/or its metabolites (6-O-monoacetylmorphine, morphine and desmethylmorphine). The two urine samples negative for heroin showed positive detection of cough medicine and methadone. One of the two urine samples also detected midazolam and zopiclone.

(Note: As morphine can also be a metabolite of codeine, detection of morphine will be counted as positive findings for opiates (heroin) if the signal of morphine detected was greater than that of codeine. In that case, the detection of codeine and codeine metabolites will not be counted as positive findings for cough medicine.)

- *Of 82 subjects who reported heroin in their last use, 30 have negative urine finding for opiates. Of the 82 subjects, 14 subjects reported last use of greater than one year, 16 subjects reported greater than one month but less than one year, 8 subjected reported greater than one week but less than one month and 1 subject did not specified the time of last use. Of the 30 urine samples, 14 of which had no drug or only therapeutic drugs detected, 5 samples were positive for methadone, 7 samples were positive for cough medicine and few samples also detected benzodiazepines, amphetamines and hypnotics.
- *Seventy-one (15.6%) of 454 urine samples have positive findings of opiates. Nineteen subjects (4.2% of total; 26.8% of 71 subjects with positive urine findings) who were under-reporting their opiates abuse in their last use were discovered by the positive urine findings.

(6) Benzodiazepines

All subjects

| History of last use | Urine detection by UPLC-TOF-MS | | |
|---------------------|--------------------------------|----------|-------------|
| | Positive | Negative | Grand Total |
| Positive | 9 | 16 | 25 |
| Negative | 52 | 377 | 429 |
| Grand Total | 61 | 393 | 454 |

*A + D = 386 (Concordance rate = 85%)

Subjects reported last drug use of < one week</p>

| History of last use | Urine detection by UPLC-TOF-MS | | |
|---------------------|--------------------------------|----------|-------------|
| | Positive | Negative | Grand Total |
| Positive | 7 | 2 | 9 |
| Negative | 26 | 192 | 218 |
| Grand Total | 33 | 194 | 227 |

*A + D = 199 (Concordance rate = 87.7%)

*If one week is taken as the window period of benzodiazepines detection in urine, the positive detection rate by UPLC-TOF-MS in subjects with positive history of benzodiazepines use is 77.8% (7 of 9 subjects). However, the number of subjects with positive urine detection and with positive history of recent use of benzodiazepines of one week or less is too small for any conclusive remark. Of the 25 subjects who reported benzodiazepines in their last use, 9 subjects reported their last use within one week's time. Two of the 9 urine samples were negative for benzodiazepines but were positive for ketamine, cough medicine and opiates.

- *Of 25 subjects who reported benzodiazepines in their last use, 16 have negative urine finding for benzodiazepines. Of the 16 subjects, 14 reported last use of greater than one week. Seven of these 14 urine samples were positive for opiates (in 4 samples), cough medicine (in 3 samples) and amphetamines (in 1 sample).
- *Sixty-one (13.4%) of 454 urine samples have positive findings of benzodiazepines. Fifty-two subjects (11.5% of total; 85.2% of 61 subjects with positive urine findings) who were under-reporting their benzodiazepines use in their last use were discovered by the positive urine findings.
- *Note: Benzodiazepines could be prescribed for therapeutic use. The number of urine samples with positive benzodiazepines was reported irrespective of the intention use as limited information of therapeutic drug history could be obtained from the study subjects.

3 Discussion

Our study recruited 454 subjects from the substance abuse clinics or non-government organizations in NTEC or NTWC. About 50 to 60 patients are arranged for follow up in each clinic session but the default rate could be high up to 30%. Our recruitment rate ranged from 0% to 33% per clinic session but in average it was less than 5%. More male (275 subjects, 60.6% of total) were recruited when comparing with female (179 subjects, 39.4% of total).

Age of study subjects ranged from 14 to 66 years. Most of the study subjects are in the younger age group of 20-29 years (118 subjects, 26% of total) and 30-39 years (159 subjects, 35% of total). However, a significant number of subject recruited (50 subjects, 11% of total) is in the teenage group. The age distribution of the study subjects has shown a prevalence of drug abusers in the younger age groups signifying a serious social problem in our locality.

Our data showed that most subjects (360 subjects, 79.3% of total) in our study population attained secondary education level, in which most of them (249 subjects, 69.1% of 360 subjects) reached junior secondary education level only. Of 454 subjects, 362 (79.7%) reported their age of first started drug abuse were below 25 years of age (Of which, 90 subjects, 19.8% of total, in 10-14 years; 201 subjects, 44.3% of total, in 15-19 years; and 71 subjects, 15.6% of total, in 20-24 years). These figures showed the importance of primary prevention of drug abuse targeting on the school-age teenagers and children (the lowest reported age of first started drug abuse was 10 years old).

In addition, long duration of drug abuse among our study subjects is observed. The longer the duration of drug abuse, the more the difficulty to achieve complete abstinence of drugs and the more the damaging effects to the subjects would be expected. These would project to a tremendously increase in the need of psychiatric services in terms of acute treatment as well as the long-term rehabilitation for substance abusers.

Irrespective of the education level attained by our study subjects, a very high unemployment rate (287 subjects, 63.2% of total) was noted in the study population.

This rate is of 18-fold higher than the unemployment rate of the general population quoted from the latest labor force statistics released by the government. Among the unemployed group, 53.4% and 63.5% of subjects are in the 20-29 years and 30-39 years age groups which should be the major workforce in the society.

Out of the total, 314 (69.2%) subjects reported to have no monthly personal income. Two hundred and nine of the 314 subjects were currently receiving financial assistance from the government. Of the 314 subjects, 161 (51.3%) reported their monthly expenditure on drugs ranging from less than \$100 to greater than \$1,100. One hundred and eleven (68.9%) of these 161 subjects were receiving financial assistance from the government. Among the 132 subjects who reported to have monthly income, 104 (78.8%) % of them have monthly income of less than or equal to \$11,000 (34.1% of these 132 subjects even have monthly income below \$5,000). Of the 132 subjects, 72 (54.5%) reported to have expenditure on drugs. Forty-one (56.9%) of these 72 subjects reported their monthly expenditure on drugs of greater than >\$1,100.

These data suggested that high unemployment rate among the drug abusers would pose a significant financial burden to the society in terms of supporting their daily-living as well as their healthcare need or even part of their expenditure on continuous drug use. Measures to enhance the employment status and monthly income of the drug abusers may improve their self-esteem and personal satisfactoriness and these may reduce their attempt of continuation of drug abuse behavior for relieving their stress. Among the employed group, continuous psychosocial counseling on tackling stress from the working environment may also be important and helpful in reducing their potential of drug use. Surveillance of continuous drug abuse behavior particularly in the group receiving social security allowance is also an important measure to promote their abstinence of drugs and as a safeguard mechanism to justify the use of our public financial resources.

Concerning the pattern of drug of abuse in our study group, discrepancy of number of drug items reported and number of drug items detected in urine was noted. Most of the subjects actually under-reported their number of drug use which can be revealed by their urine analysis results. These data may suggest that performing survey or monitoring of drug use behavior in the format of questionnaire would under-estimate the actual situation of the drug abuse problem. On the contrary, analysis of biological samples could provide an additional objective evidence of individual's drug of abuse practice and this could be helpful in the surveillance of the drug of abuse problem globally for planning of the drug-beating strategy and individually for formulating a better clinical management.

Drug abuse is regarded as a private matter that most people would like to hide away from others by under-reporting the issue. Even though urine analysis may have limitations in the window period of detection of different drugs or drug abusers can adulterate their urine samples to create false negative results, different measures can be implemented to ensure the reliability of the sample collection and to enhance the sensitivity and specificity of drug detection in biological samples. Our data showed that the number of drug items detected in urine was much higher than that reported voluntarily by the subjects. Besides, urine analysis could help to monitor the absolute abstinence of drug abuse and to detect any change of pattern of drug abuse. More recently, hair analysis for detection of chronic drug abuse is of a hot-topic issue under evaluation of its potential clinical use by multiple centers worldwide. By using a biological sample that could allow us to detect the metabolites of abused drug remaining in the body, a longer window of detection could be achieved and a higher drug detection rate would be obtained in those who have negative urine findings.

UPLC-TOF-MS is one of the state of the art technology commonly used in toxicology screening and confirmation based on a set of stringent criteria of matching of the accurate mass and fragmentation pattern of the molecule with the data of the drug library. For the comparison of the efficacy of drug detection by the new (UPLC-TOF-MS) and conventional (HPLC-UV and Immunoassay) techniques, our current data showed that the new technique is able to detect more drugs than the conventional techniques. Our finding together with the previous published data [HK Lee *et al.* Anal Chim Acta (2009) 649:80-90] supported the use of UPLC-TOF-MS for clinical urine toxicology services. Applications of this technology in other matrices, such as oral fluid and hair, shown promising by published data from literature and preliminary data of our group, require further evaluation.

Regarding to the findings of our urine analysis, cough medicine is the most frequently drug item detected in our study population. Several reasons could be postulated from our findings. The easy accessibility from local pharmacy and cheaper cost of cough medicine could be the reasons for the use of cough medicine by chronic drug abusers as a milder substitute of their original drug use or during their detoxification process.

Besides, the unexpected finding of cough medicine in our study subjects who did not reported history of cough medicine use may suggest that ingredients of cough medicine may be frequently added to the commonly abused drugs. Whether the addition of cough medicine is intentional for the synergistic sedative effect or for minimizing some of the unwanted effects of the abused drugs, or unintentional as contaminants, further study and investigation is required. From our observation, 10 of the 74 urine samples positive for ketamine also had positive findings of cough medicine (such promethazine, chlorpheniramine, brompheniramine, as pseudoephedrine/ephedrine, dextromethorphan, propoxyphene, codeine, cetirizine and diphenhydramine). Ketamine is a well-known dissociative anaesthetics with short half-life of 2.5 to 3 hours. Promethazine, one of the most commonly detected cough medicine ingredient in our study population, is an anti-histamine with potent sedative effect and also anti-emetic, anti-motion-sickness and anti-cholinergic effects. Clinically, the combined use of ketamine and promethazine has been described in paediatric dental use for sedative purpose as promethazine can help to reduce the emetic side-effect from ketamine while the sedative effects are good.

Although cough mixture has been noted as an emerging psychiatric problem in Hong Kong since 1990s, the potential toxic effects of cough medicine are unnoticed by the general public. Previous local study showed that acute organic brain syndrome, schizophreniform psychosis and affective episode were the main psychiatric presentations which appeared to be associated with the pharmacological activities of opiates, antihistamines and sympathomimetics, the main ingredients of most cough mixtures. Despite psychiatric complications caused by cough medicine, metabolic changes, such as severe hypokalaemia and metabolic acidosis that potentially leads to fatal arrhythmia, had been reported in the literature. A local study also demonstrated cough mixture abuse as a novel cause of folate deficiency and reported a case of cerebellar degeneration and folate deficiency due to cough mixture abuse.

Social pressure from peers, family problems, ease of access, and unawareness of the consequence of cough medicine misuse had been reported as the primary factors accounting for adolescent cough medicine abuse. Our findings provide further evidence to the commonness of the cough medicine use among our youngsters which is currently under-reported and under-recognized in our locality.

Non-benzodiazepine hypnotics such as zopiclone and zolpidem were of growing popularity in our locality which were also observed in our findings. Zopiclone, a benzodiazepine-like drug with half-life of about 6 hours, believed to have less dependence problem than benzodiazepines, is in fact having a greater additive potential than traditional benzodiazepines. Zopiclone is usually recommended for short-term use. However, because of the easy accessibility of non-benzodiazepines hypnotics from local pharmacy and the ignorance to the potential dependence and withdrawal symptoms with long-term use, many people who have insomnia due to stress or night-shift duties would buy these drugs over-the-counter and become chronic users of hypnotics. Chronic zopiclone users reported withdrawal symptoms like anxiety, tachycardia, sweating, palpitations and tremor frequently and also further insomnia despite the use of high dosage of hypnotics. Unusual haematological complications like haemolytic anaemia and methaemoglobinaemia had been reported locally. Delirium and convulsions due to withdrawal of zopiclone, and fatal cases due to severe zopiclone overdose have been reported in the The damaging effects of chronic hypnotics misuse are seriously literature. under-recognized in our locality.

Ketamine has remained the most common psychotropic substance of abuse among youngster in Hong Kong since 2001. According to the data reported by the Central Registry of Drug Abuse (CRDA), 85.4% of the reported young drug abusers aged under 21 abused ketamine in 2008 and the latest figure was 78.2% in 2010. The increasing concern of ketamine use in the society is not only because of its growing popularity among the teenagers, but also a significant rising trend of traffic accidents caused by driving under the influence of drugs (DUID). Driving under the effect of ketamine would significantly impair the driving performance putting the drug drivers at an unrecognized risk of personal harm as well as a tremendous danger to other road users.

According to the data collected by the Hospital Authority Hong Kong Poison Information Centre (HKPIC), ketamine abusers represented 16% of all drug abusers attending accident and emergency departments (AEDs) in the period of 1 July 2005 to 31 December 2005, and the proportion rose significantly to 40% in the period of 1 January 2008 to 30 June 2008. In a retrospective review of the acute clinical presentations of ketamine abusers in fifteen accident and emergency departments in Hong Kong, most ketamine abusers in the series were young (84% being 13-29 years old), male, and presented with impaired consciousness, abdominal pain, or dizziness. In their series of ketamine users, up to 21% and 16% of cases presented

with abdominal pain and abnormal liver function test results respectively. In another local study, ketamine abusers were noted to have upper gastrointestinal symptoms frequently, the commonest of which is epigastric pain.

In our study, 113 subjects reported ketamine in their last drug use, of which, 80 of them did not specify any physical complication. Seven (21.2%) of the 33 subjects who had physical complications reported epigastric pain as one of their complaints. Six of these seven ketamine abusers also complained urinary problems. These subjects are young, aged between 23 to 36 years, 4 are males and 3 are females.

Recent local observations also suggested that long-term ketamine usage can cause biliary abnormalities resulting in recurrent epigastric pain and elevated ductal enzymes. In a case report, the maximal diameter of the dilated common bile duct can be up to 17mm which is about five- to six-fold of the usual dimension (about 3mm) in normal young adults and can be misdiagnosed as another biliary tree abnormality, namely choledochal cyst, that may lead to unnecessary investigations or even an operation. Therefore, apart from the growing alertness of the ketamine-induced cystitis and urinary tract problems, the general public should be made aware of this ketamine-related epigastric/abdominal pain and hepatobiliary problems.

Similar drug of abuse items were detected in the subgroups of NTEC and NTWC, differing by the order of frequency. Cough medicine and hypnotics were the two most commonly detected drug items in our study population. This pattern is different from the usual pattern reported by the CRDA. The physical and psychological dependence, as well as the psychiatric complications caused by cough medicine and hypnotics that are commonly encountered in patients of SACs are currently overlooked by the community. Education to the general public on the harmful effects of these two benign-looking therapeutic medications would be helpful in reducing the number of new cough medicine and hypnotics abusers who try them by ignorance. More stringent policy on monitoring local pharmacies on the selling of unprescribed cough medicine and hypnotics could also be considered by the government.

4 Conclusions

Our report has presented the findings of the survey and urine analysis conducted in 454 subjects from the substance abuse clinics or non-government organizations in NTEC or NTWC during the period of July 2007 to March 2011.

The demographic characteristics of our study population with their continuing substance abuse habit and long-term psychiatric problems created a huge social, financial and healthcare burden to the society. While primary prevention of substance abuse is of paramount importance to avoid these potential problems from happening in all generations, effort and resources on tackling the existing drug abusers in terms of stopping and monitoring their drug misuse habit, treating their physical and psychiatric illnesses, and ultimately helping them to return to the society with public acceptance, cannot be neglected.

Differences in age revealed a different prevalence of drug abuse pattern in the locality. Though the most commonly detected drug of abuse items in NTEC and NTWC were similar, the differences in the frequency of detection of individual drug items can be explained by the different age components among the regional subgroups. More young subjects in NTEC showed a higher prevalence of ketamine abuse while more old subjects in NTWC showed a higher prevalence of opiates abuse. Nevertheless, the commonness of cough medicine (26.7% of total) and hypnotics (18.3% of total) misuse among patients followed up in SACs across the new territories gives an important message to the community that their potential harmful effects should not be overlooked.

Advanced technology in drug testing is useful in diagnosing and monitoring substance abuse habit, as well as in discovering unusual pattern and novel or contaminated substances being misused, though cost and window of detection would be its limitations.

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7 Appendix

7.1 Questionnaire

| Part I: Demographic Data | |
|------------------------------------------------------------------------------------------|-----|
| Sex \Box Male / \Box Female | |
| Age or DOB year-old or/ / (DD/MM/YY) | |
| Marital Status Single / Co-habitant / Married / Separated / Divorced / Widow | ved |
| Educational Attainment | |
| \Box No formal education | |
| □ Pre-Primary education | |
| \Box Primary education (Complete / Incomplete: up to Primary) | |
| \Box Secondary education (Complete / Incomplete: up to Form) | |
| □ Tertiary education: IVE / Diploma / Higher-diploma / Associates / University | |
| □ Others (please specify:) | |
| Current Employment | |
| □ Occupation: (□Full-time / □Part-time; For weeks / months / years) | |
| □ Unemployed (For weeks / months / years, previously worked as:) | |
| □ Student (□Full-time / □Part-time, please specify:) | |
| Monthly Personal Income | |
| \Box No income (Did you receive any social security allowance? \Box Yes / \Box No) | |
| \Box Less than \$1,000 | |
| □ \$1,000 to \$2,999 | |
| □ \$3,000 to \$4,999 | |
| □ \$5,000 to \$6,999 | |
| □ \$7,000 to \$8,999 | |
| □ \$9,000 to \$10,999 | |
| □ >\$11,000 | |
| Monthly Expenditure on Drugs | |
| \Box Nil \Box Less than \$100 | |
| □ \$100 to \$299 | |
| □ \$300 to \$499 | |
| □ \$500 to \$699 | |
| □ \$700 to \$899 | |
| □ \$900 to \$1,099 | |
| □ >\$1,100 | |

Part II: Information on Drug Use

Is this your first visit to CPH Substance Abuse Clinic? \Box Yes / \Box No If No, when did you first visit this clinic? _____ (MM/YY)

When did you first try the drug(s)?

Age / Year ____

What drugs have you tried?

Narcotics Analgesics

□ Opiates / Heroin (海洛英/白粉)

(Mode of Administration: oral / smoking / chasing dragon / iv injection)

Tranquillizers

□ Benzodiazepine / Midazolam (Dormicum 藍精靈) / Diazepam (Valium) / Others

□ Hypnotics / Zopiclone (白瓜子)

Stimulants

□ Amphetamine / Methylamphetamine (冰)

□ MDMA (Ecstasy / 搖頭丸)

□ Cocaine (可卡因)

Hallucinogens

□ Cannabis (大麻 / 草)

 \Box Cannabis resin

 \Box LSD

Depressants

□ Methaqualone (Mandrax 忽得)

Others

- □ Ketamine
- \Box Cough Medicine
- □ Thinner / Organic Solvent

 \Box Others, please specify:

 \Box Not known

Why did you try the drug(s)?

Experimental / Recreational / Dependence / Peer Influence / Others (Please specify: _____)

_____;

Did you suffer from any complications?

□ Yes, (Physical: _____

Psychiatric: Psychosis / Depression / Impaired memory / Others: _____)

 \Box No

When did you take the drugs lastly?

Exact Date / / (DD/MM/YY) Estimated Time _____ hours / days / weeks / months / years ago

What drugs have you taken lastly?

Please specify:

Where and how did you get the drug(s)?

Where did you take the drug(s)?

| Disco / Bar / Karaoke / Video Game Centre / Pharmacy / Home | / Friend's Apartment / Street |
|-------------------------------------------------------------|-------------------------------|
| Others, please specify: | |

Are you taking any regular medications now?

| \Box Yes. (Please specify: |) |) |
|------------------------------|---|---|
| □ No | | |

Smoking and Drinking History:

| □ Smoker | □ Ex-Smoker | □ Non-Smoker | |
|------------|----------------|--------------|---------------|
| Drinker (R | egular/Social) | □ Ex-Drinker | □ Non-Drinker |

Forensic History:

🗆 Nil

| □ Yes (Please specify: rol | obery / theft / assault / | / drug trafficking / dru | ug possession / others) |
|----------------------------|---------------------------|--------------------------|-------------------------|
| Currently on Probation: | \Box Yes \Box No | | |

~ End of Questionnaire ~