

# Portable Electrochemical Detection of Illicit Drugs in Smuggled Samples: Towards More Secure Borders <sup>†</sup>

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**Abstract:** Illicit drug consumption is posing critical concerns in our society causing health issues, crime-related activities, and the disruption of the border trade. The smuggling of illicit drugs urges the development of new tools for rapid on-site identification in cargos. Current methods used by law enforcement offices rely on presumptive color tests and portable spectroscopic techniques. However, these methods sometimes exhibit inaccurate results due to commonly used cutting agents or because the drugs are smuggled (hidden or mixed) in colored samples. Interestingly, electrochemical sensors can deal with these specific problems. Herein, it is presented an electrochemical device that uses low-cost screen-printed electrodes for the electrochemical detection of illicit drugs by square-wave voltammetry (SWV) profiling. A library of electrochemical profiles is built upon pure and mixtures of illicit drugs with common cutting agents. This library allows the design of a tailor-made script that shows the identification of each drug through a user-friendly interface. Finally, the results obtained from the analysis of different samples from confiscated cargos at different end-users sites present a promising alternative to current methods offering low-cost and rapid testing in the field.

**Keywords:** Electrochemical sensors; square-wave voltammetry; screen-printed electrodes; illicit drugs; forensics; portable device

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## 1. Introduction

The consumption of drugs of abuse is causing critical issues in our society due to health issues, crime-related activities, and the disruption of the border trade [1]. These illicit drugs can enter the illegal market through external borders (e.g. natural drugs) or by internal production (e.g. synthetic drugs). The smuggling of illicit drugs such as cocaine and heroin in Europe urges the development of new tools for rapid on-site identification in cargos. Besides, the production of synthetic drugs increases internal trafficking, thus demanding simple and straightforward devices to detect illicit drugs in the field. Current methods used by law enforcement offices rely on presumptive color tests [2] and portable spectroscopic techniques (e.g. near-infrared [3] and Raman spectroscopy [4]). However, these methods sometimes exhibit inaccurate results due to commonly used cutting agents or because the drugs are colored. Besides, drug traffickers are generating innovative ways to overcome traditional detection methods such as mixing with conventional goods (e.g. charcoal, food) or adding colorants or other substances to avoid the on-

site determination by current methods. Therefore, new devices that can overcome the current problems are necessary to cope with the determination of smuggled illicit drugs in common goods.

The devices for on-site analysis must be portable, low-cost and user-friendly in order to be implemented and used by law enforcement officers [5]. Electrochemical sensors can provide the aforementioned features, and importantly, they can deal with current challenges, providing more reliable results in comparison to commercially available devices [6,7]. In this direction, portable and wearable electrochemical sensors have been designed for the detection of illicit drugs in different configurations [8], including glove-based sensors [9]. The electrochemical approach is based on the characteristic electrochemical profile (EP) of each compound that reveals the electroactive moieties of the target compound. Following this strategy, cocaine [10], ketamine [11], heroin [12] and synthetic cathinones [13] have been detected by using low-cost screen-printed electrodes (SPEs). Amphetamine is a special case as it is not electroactive in the potential window of commercial carbon SPEs. Therefore, an in situ derivatization by employing 1,2-Naphthoquinone-4-sulphonic acid sodium salt allows for its electrochemical detection [14]. Overall, the most used illicit drugs can be determined by electrochemical methods at certain conditions.

Herein, it is presented an electrochemical device that uses low-cost SPEs for the electrochemical detection of illicit drugs by square-wave voltammetry (SWV) profiling. A pH strategy based on the profiling of each illicit drug using specific buffers allows detecting the most encountered illicit drugs (i.e. cocaine, MDMA, heroin, and amphetamine). Hence, the electrochemical interrogation of the illicit drugs exhibits the oxidation of the electroactive moieties in each drug at a certain potential, with the exception of amphetamine that uses an in situ derivatization to unravel its oxidation peak. A library of electrochemical profiles is built upon pure and mixtures of illicit drugs with common cutting agents. This library allows the design of a tailor-made script that shows the identification of each drug through a user-friendly interface. Finally, the results obtained from the analysis of different samples from confiscated cargos at different end-users sites present a promising alternative to current methods. Overall, the fast analysis of samples with a portable electrochemical device exhibits a straightforward on-site detection aiming to facilitate the tasks of law enforcement agents in the field, thus providing a more secure border management and a safer society.

## 2. Methods

### 2.1. Materials

Standards of D,L-amphetamine · HCl, methamphetamine · HCl, 3,4-methylenedioxymethamphetamine · HCl (MDMA), cocaine · HCl and heroin · HCl, were purchased from Chiron AS, Norway. Standards of paracetamol, caffeine and creatine were provided by National Institute for Criminalistics and Criminology (NICC, Belgium). Confiscated samples of amphetamine, MDMA, cocaine and heroin were also provided by the NICC. Analytical grade salts of potassium chloride, potassium phosphate, sodium borate, sodium bicarbonate, sodium acetate and potassium hydroxide were purchased from Sigma-Aldrich (Overijse, Belgium). 1,2-naphthoquinone-4-sulphonic acid sodium salt (NQS) (>98%) was purchased from Tokyo Chemical Industry Co., LTD., Japan.

### 2.2. Methods

Square wave voltammograms and cyclic voltammograms were recorded using a MultiPalmSens4 or EmStat Pico potentiostats (PalmSens, The Netherlands) with PSTrace/MultiTrace. Disposable ItalSens SPEs (PalmSens, the Netherlands), containing a graphite working electrode ( $\varnothing = 3$  mm), a carbon counter electrode, and a (pseudo) silver reference electrode were used for all measurements. The SWV parameters that were used: potential range of 0.0–1.4 V, frequency 10 Hz, 25 mV amplitude and 5 mV step potential.

All the square wave voltammograms are background corrected using the PStTrace software.

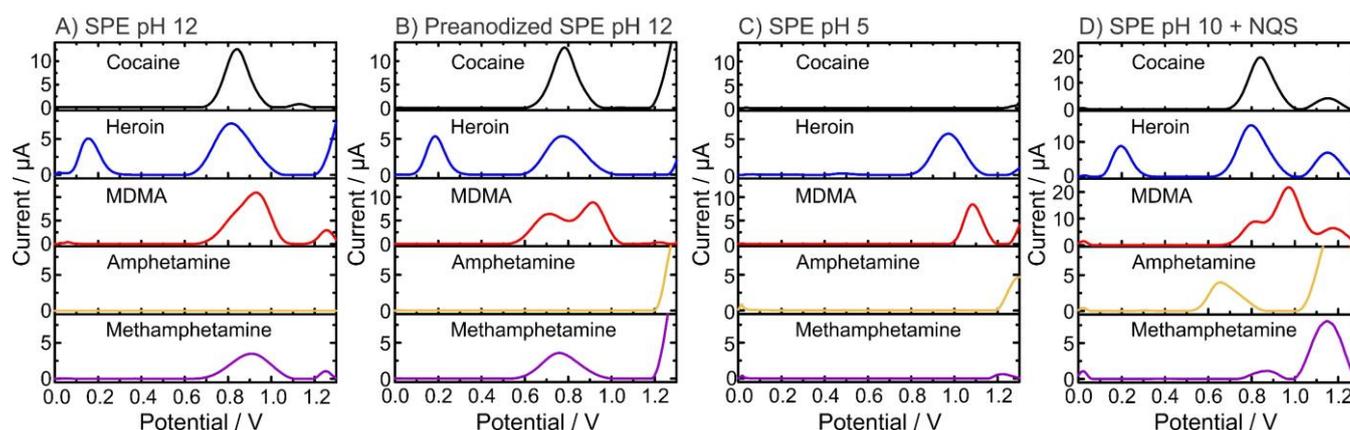
Electrochemical tests were performed in 20 mM buffer solutions with 100 mM KCl by applying 60  $\mu$ L of the solution onto the SPE. Phosphate buffer, acetate buffer and hydrogen carbonate buffer were used for the detection of cocaine and heroin, MDMA and amphetamine, respectively. Preanodized SPEs for heroin detection were performed by applying 1.5 V for 60 s in PBS solution at pH 7 by drop casting 60  $\mu$ L on the SPE [12].

The composition of the confiscated samples was previously analyzed in the forensic laboratory at NICC with gas chromatography-mass spectrometry (GC-MS) to subsequently validate the electrochemical approach. Besides, the confiscated samples were also analyzed by a handheld Raman spectrometer (Bruker Bravo, Germany).

### 3. Results and Discussion

#### 3.1. Electrochemical Profiling of Illicit Drugs

The electrochemical profiling is based on the interrogation by SWV of the target molecules which exhibit an oxidation process at certain potential. Hence, cocaine oxidation peak might differ from heroin oxidation peak, showing the possibility to identify the target by the specific peak potential [15]. Fortunately, illicit drugs contain moieties which are electroactive at the potential window of carbon SPEs. These moieties are usually secondary or tertiary amines that allows its oxidation, thus showing a peak signal during the electrochemical scan by SWV. However, some illicit drugs share some moieties which can exhibit some overlap in the oxidation potential. In previous works, our research group has optimized the detection of illicit drugs by exploring certain conditions during the SWVs. For example, the anodic pretreatment to unravel the phenolic group oxidation of the 6-monoacetylmorphine (6-MAM) (a byproduct of heroin degradation at pH 12) [12], the cathodic pretreatment to avoid the suppressing effect of some adulterants [13] or the use of different pH (as some moieties are not oxidizable in certain pH at SPE) [11]. Besides, some illicit drugs such as amphetamine need a derivatization step to allow its electrochemical profiling employing low-cost carbon SPEs. In this case, a simple mixing step with NQS launches a chemical reaction to a product that is electroactive at the SPE [14].



**Figure 1.** Electrochemical profiles of illicit drugs (0.5 mM) obtained by square-wave voltammetry (SWV) using SPE at different pH: **A)** pH 12; **B)** pH 12 using preanodized SPE; **C)** pH 5; and **D)** pH 10 including the derivatizing agent NQS.

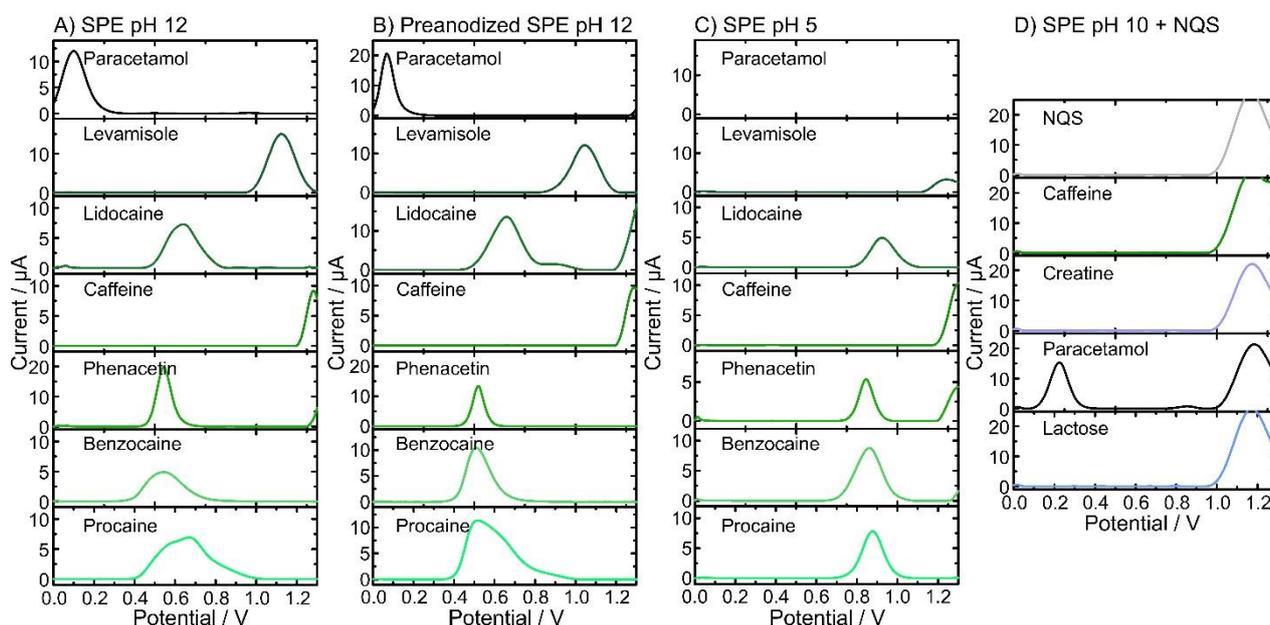
#### 3.2. Generating the Library of Electrochemical Profiles

Figure 1 shows the SWVs of pure illicit drugs (i.e. cocaine, heroin, MDMA and amphetamine) at pH 5, pH 10, pH 12 and pH 12 using anodic pretreated SPEs to record the specific electrochemical profiles of the molecules at certain conditions. Figure 1A and Figure 1B shows the differences between anodic pretreated SPEs. Although similar profiles were obtained, a clear peak separation occurred in the MDMA signal. The pH dependence

on the oxidation of illicit drugs is dramatically shown in the analysis at pH 5, where only heroin and MDMA exhibited electroactivity (Figure 1C). This fact assists on the proper identification of the unknown sample by a simple dual pH test. Finally, the pH assessment clearly exhibits that amphetamine is not electroactive, and only after the derivatization step (Figure 1D), an oxidation peak appears.

A similar electrochemical approach was performed employing the most encountered cutting agents (i.e. paracetamol, levamisole, lidocaine, caffeine, phenacetin, benzocaine, procaine and lactose [12,14,16]) (Figure 2A-D). Particularly interesting is the effect of the anodic pretreatment on the paracetamol signal exhibiting a sharper oxidation peak (Figure 2B). This permits the proper identification of the 6-MAM peak, thus avoiding peak overlap [12]. Besides, the effect of pH on the electrochemical signal showed a pH dependence as the oxidation peak shifts towards higher potentials at acidic pHs (Figure 2C). As pH 10 with NQS is targeted for the detection of amphetamine, only common cutting agents encountered in amphetamine real samples are explored (Figure 2D). Considering the profiling of the cutting agents, most of the peak potentials do not fall in the same position as the illicit drugs, thus allowing for a suitable identification in real samples.

After building the library of electrochemical profiles with several conditions, a custom-made script (Matlab R2018b, MathWorks, USA) is integrated. This script enhances the peak separation and facilitates the identification of the compounds in the sample. In brief, the script removes the background signal and applies a top-hat filter that provides an enhanced separation of overlapped peaks which permits a successful identification of the substances based on the peak potential of each drug. Therefore, the peak potential of each drug and cutting agent is introduced in the script to properly identify the drug and display it through a user-friendly interface.

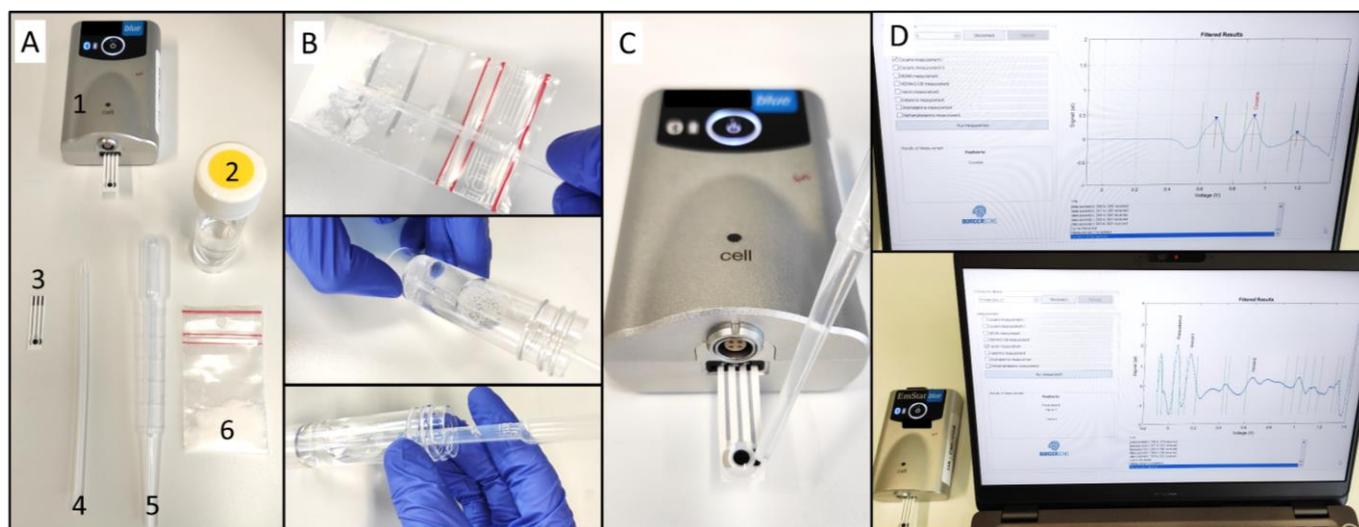


**Figure 2.** Electrochemical profiles of common cutting agents (0.5 mM) obtained by square-wave voltammetry (SWV) using SPE at different pH: **A)** pH 12; **B)** pH 12 using preanodized SPE; **C)** pH 5; and **D)** pH 10 including the derivatizing agent NQS.

### 3.3. Testing the Portable Electrochemical Device with Confiscated Samples

The electrochemical device consists of a miniaturized potentiostat with Bluetooth connectivity, a disposable SPE, a sampling container, a disposable spatula and a disposable pipette (Figure 3A). The sampling procedure consists of collecting the powder (either powder, liquid, crystal or impregnated material) with the disposable spatula into a tube containing 15 mL of the suitable buffer (Figure 2B). After shaking thoroughly, a drop of

the solution is deposited on the SPE with the disposable pipette (Figure 2C). Subsequently, the operation is started on the user-friendly interface launching the electrochemical method, subsequent data treatment and results display (Figure 2D). For the analysis of confiscated samples, the strategies employing pH 12, preanodized SPE in pH 12, pH 5 and pH 10 with NQS were employed for cocaine, heroin, MDMA and amphetamine, respectively.



**Figure 3.** On-site detection of illicit drugs with the portable electrochemical device. **A)** Elements of the electrochemical device (1-potentiostat, 2-buffer container, 3-SPE, 4-disposable spatula, 5-disposable pipette, 6-confiscated sample); **B)** Sampling procedure; **C)** deposition of the solution on the setup ready for the electrochemical interrogation; and **D)** user-friendly interface showing the results of the analysis.

The reliability of the electrochemical device was evaluated in 40 confiscated samples provided by NICC (Table 1). After the analysis, the 40 samples were all positive for the corresponding illicit drug using the described sampling method in comparison to the standard methods (GC-MS). Besides, a portable Raman spectrometer was also used as a commonly used method in border settings. The electrochemical reader and portable Raman spectrometer exhibited an accuracy of 100% and 50%, respectively. Therefore, the electrochemical device outperformed the Raman device, particularly in the heroin and amphetamine detection. It is worth to mention that the low performance of the Raman device could be attributed to the colored nature of the samples, thus exhibiting one of the flaws of current methods. Overall, the electrochemical device is positioned as a reliable alternative for its use in the field due to its affordability, reliability and user-friendliness.

**Table 1.** Results of the analysis by the analytical methods and composition of the confiscated samples.

Sample	GC-MS	Electrochemical reader	Portable Raman
1	Cocaine	Cocaine	Cocaine mixture
2	Cocaine, caffeine, levamisole	Cocaine	Cocaine mixture
3	Cocaine, levamisole	Cocaine	Cocaine mixture
4	Cocaine, lidocaine, levamisole, phenacetin	Cocaine, lidocaine	Lidocaine
5	Cocaine, phenacetin, levamisole	Cocaine, phenacetin	Cocaine mixture
6	Cocaine, levamisole	Cocaine	Cocaine mixture
7	Cocaine base	Cocaine	Benzyl benzoate
8	Cocaine, levamisole	Cocaine, levamisole	Cocaine mixture
9	Cocaine, caffeine	Cocaine	Cocaine mixture
10	Cocaine, lidocaine, levamisole, caffeine, phenacetin	Cocaine, lidocaine	Lidocaine
11	Heroin, caffeine, 6-MAM, Noscapine, Papaverine	Heroin	Normorphine
12	Heroin, caffeine, 6-MAM, Morphine, Noscapine, Papaverine	Heroin	Cetyltrimethylammonium chloride
13	Heroin, caffeine, paracetamol, 6-MAM, Noscapine, Papaverine	Heroin, paracetamol	Cetyltrimethylammonium chloride
14	Heroin, caffeine, Dextromethorphan, 6-MAM, Noscapine	Heroin	Unknown
15	Heroin, 6-MAM, Noscapine, Papaverine	Heroin	Cetyltrimethylammonium chloride
16	Heroin, paracetamol, 6-MAM, morphine, caffeine, codeine, noscapine, papaverine	Heroin, paracetamol	Cetyltrimethylammonium chloride
17	Heroin, paracetamol, 6-MAM, caffeine, noscapine, papaverine	Heroin, paracetamol	Cetyltrimethylammonium chloride
18	Heroin, paracetamol, 6-MAM, caffeine, noscapine, papaverine	Heroin	Unknown
19	Heroin, paracetamol, 6-MAM, caffeine, noscapine, papaverine	Heroin, paracetamol	Cetyltrimethylammonium chloride
20	Heroin, paracetamol, 6-MAM, caffeine, noscapine, papaverine	Heroin, paracetamol	Cetyltrimethylammonium chloride
21	MDMA	MDMA	MDMA tablet
22	MDMA	MDMA	MDMA crystals
23	MDMA	MDMA	MDMA tablet
24	MDMA, levamisole	MDMA	MDMA crystals
25	MDMA	MDMA	MDMA tablet
26	MDMA	MDMA	MDMA tablet
27	MDMA	MDMA	MDMA crystals
28	MDMA	MDMA	MDMA crystals
29	MDMA	MDMA	MDMA crystals
30	MDMA	MDMA	MDMA crystals
31	Amphetamine	Amphetamine	Paraform
32	Amphetamine	Amphetamine	Norephedrine HCl
33	Amphetamine	Amphetamine	1_(S)-2-diphenylmethylpyrrolidine HCl_(S)-desoxy-D2PM
34	Amphetamine	Amphetamine	1_(S)-2-diphenylmethylpyrrolidine HCl_(S)-desoxy-D2PM

35	Amphetamine, caffeine	Amphetamine	Amphetamine
36	Amphetamine, caffeine	Amphetamine	Unknown
37	Amphetamine	Amphetamine	1-phenyl-1-propanol
38	Amphetamine	Amphetamine	1-phenyl-1-propanol
39	Amphetamine	Amphetamine	1-phenyl-1-propanol
40	Amphetamine, caffeine	Amphetamine	Amphetamine

#### 4. Conclusions

In this work, the analysis of confiscated samples from illicit drugs is presented by the use of a portable electrochemical device. First, the construction of a library from several electrochemical profiles of standards of illicit drugs and common cutting agents at different conditions by SPE is performed. After the selection of the suitable conditions and the integration of the peak potentials of each target into a tailor-made script, the electrochemical device is ready for the on-site analysis. The examination of 40 confiscated samples with the electrochemical device and a portable Raman spectrometer showed an outstanding performance of the electrochemical device in front of the Raman device according to the GC-MS identification. Overall, the electrochemical device based on SPE is presented as a promising alternative to current rapid and on-site methods for the detection of illicit drugs at border and coast controls.

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