# Local Anaesthetics – Substances with Multiple Application in Medicine

# **Rodica SÎRBU**

Ovidius University of Constanta, Faculty of Pharmacy, Constanta, Romania

### **Emin CADAR**

UMF Carol Davila Bucharest, Faculty of Pharmacy, Bucharest, Romania

# Cezar Laurențiu TOMESCU

Ovidius University of Constanta, Faculty of Medicine, Constanta, Romania

### Cristina-Luiza ERIMIA

Ovidius University of Constanta, Faculty of Pharmacy, Constanta, Romania

# **Stelian PARIS**

Ovidius University of Constanta, Faculty of Pharmacy, Constanta, Romania

# **Aneta TOMESCU**

Ovidius University of Constanta, Faculty of Medicine, Constanta, Romania

#### Abstract

Local anaesthetics are substances which, by local action groups on the runners, cause loss of reversible a painful sensation, delimited corresponding to the application. They allow small surgery, short in duration and the endoscopic maneuvers. May be useful in soothe teething pain of short duration and in the locking of the nervous disorders in medical care. Local anesthesia is a process useful for the carrying out of surgery and of endoscopic maneuvers, to soothe teething pain in certain conditions, for depriving the temporary structures peripheral nervous control. Reversible locking of the transmission nociceptive, the set of the vegetative and with a local anesthetic at the level of the innervations peripheral nerve, roots and runners, a trunk nervous, around the components of a ganglion or coolant is cefalorahidian practice anesthesia loco-regional. Local anaesthetics summary and semi-summary have multiple applications in dentistry, consulting, surgery and obstetrics, constituting "weapons" very useful in the fight against the pain.

Keywords: Local anaesthetics, dentistry, consulting, surgery and obstetrics

### Introduction

Local anesthesia is a useful method for performing surgical procedures and endoscopic maneuvers, in order to relieve pain in certain conditions, for the temporary depravation of peripheral structures from nervous control. Reversible locking of the nociceptive, motor and vegetative transmission with a local anesthetic at the level of peripheral nerve endings, nerve roots, of a nerve trunk, around the components of a ganglion or in the cerebrospinal fluid constitutes the practice of locoregional anesthesia. Local anaesthetics are substances which, through local action on nerve formations, lead to the reversible loss of the pain sensation, defined corresponding with the application area [1]. The first known local anesthetic was cocaine. It is found in the leaves of a shrub scientifically called Erythroxylon coca, and more commonly coca shrub. The shrub grows wildly in the Andes Mountains of South America, but also in Ceylon and Jamaica. The natives use it by chewing the leaves, to remove the sensation of hunger, thirst and fatigue. An American author noted that natives measure the distance between villages in coca leaves [2]. Later, in 1948, xiline or lidocaine was introduced with a faster and more intense action than procaine and, also with a notable efficacy in the treatment of cardiac arrhythmias [1, 2]. Xiline was followed, in 1952, by chloroprocaine, which has, however a higher level of neurotoxicity, in 1957 by mepivacaine, acting somewhat longer than xiline and bupivacaine, introduced in 1963 [1,2]. The latter local anesthetic determines a longer action period and it is indicated to be used to remove the pain of childbirth or postoperative pains. Administered through continuous infusion, it may cause effective analgesia for several days. Unfortunately, its administration must be carefully monitored because it has side effects on the heart. For the anesthesia of the skin and of the mucous membranes were later synthesized other local anaesthetics. As it can be seen, cocaine, the first anesthetic discovered lost, ground to the newer anaesthetics, which is also because this preparation snorted through the nose causes a feeling of drunkenness and an increase of physical and mental energy, phenomena sought after avidly by addicts. The chronic use of cocaine with this purpose determines however phenomena of somatic damage, the chronic consumer having an earthy yellow color, with a total lack of appetite, physically and mentally exhausted, phenomena that can lead to death. Clinically, cocaine is now used less often, only for the surface anesthesia of the mucous membrane of the eyes and of the nasopharynx. However, the other local anaesthetics have multiple uses in dentistry, ENT surgery and obstetrics, very useful "weapons" in the fight against pain. However, most local anaesthetics are injected near the action area and they must be able to penetrate the membrane of the nerve. In general, the membranes of the nerves are composed of lipids. The increase of solubility of lipids of a series of compounds has as a result the facilitation of membrane penetration. In *in vitro* experiments involving very simple systems with isolated nerves, the potency of the compounds is directly proportional to the distribution coefficient. The *in vivo* system is more complicated and often, in a homologous series, the increase of partition coefficients leads to the increase of potency to a maximum, afterwards the activity decreases, at the same time the toxicity increases [1, 3].

Local anaesthetics of synthesis and semi-synthesis have multiple uses in dentistry, ENT, surgery and obstetrics, constituting very useful "weapons" against pain.

# In this paper the local anaesthetics are systematized depending on their structure. Assessments are made regarding the effects based on the correlations generated through the analysis of the structures from spectroscopy data [3, 4].

# **Research Methods**

In this study the information regarding the natural and synthetic anaesthetics is systematized, with the presentation of the chemical structures, short characterizations and highlights of the beneficial and toxic effects.

Through the spectrophotometric method in infrared were obtained spectra in infrared for the following local anaesthetics: Cocaine, Prilocaine, Lidocaine, Bupivacaine, Benzocaine, Procaine. The FT IR 4200 Jasco spectrometer with Fourier transform, manufactured in Japan, Tokyo, was used. The characteristics of the device are: the wavelength 7800 to 350cm-1; with the precision of  $\pm 0.01$  cm<sup>-1</sup> and the maximum resolution of 0,5 cm<sup>-1</sup>[4]. Spectra in infrared were carried out for the following substances: Cocaine, Benzocaine, Procaine, Lidocaine, Bupivacaine, Prilocaine.

# **Results and Discussions**

### A systematization of local anaesthetics can be made after the chemical structure and the obtaining method/ In Table 1 and 2 are presented examples of local anaesthetics [3]

Local	Effect on the human body
anaesthetics and	
structure	
Cocaine	Cocaine is an alkaloid isolated from Erxtroxylon coca
	leaves, Family Erytroxylaceae, in 1855, by Gaedcke, who
COOCH3	assumed that it is an alkaloid related to caffeine. In 1860
×	Niemann obtained it by extraction from the leaves, along
H3C-N ) - 0-C-C6H5	with other compounds with a similar chemical structure:
ö	benzoylecgonine, cynnamil cocaine. The fruits have been
	used by locals since ancient times, for their effects (elation
	and removing the sensation of fatigue, hunger and thirst).
	Cocaine is obtained also synthetically. The cocaine can
	cause very serious acute poisoning (some with a fatal
	outcome), especially as therapeutic accidents (overdose,

 Table 1. Natural local anaesthetics [1,3, 5,6]

$\begin{array}{c} CH_2 - CH - \cdots - CH - CO - 0CH_3 \\   & 1 \\ N - CH_3 \\ I \\ CH_2 - CH - \cdots - CH_2 \end{array}$	confusion, or consecutive to the hypersensitivity of the subject), rarely because of suicide and chronic poisoning - cocainomania. Base cocaine is rarely used, as oil solutions 2%, used in ophthalmology. Its salts are frequently used: cocaine hydrochloride and nitrate. Cocaine does not penetrate intact skin, but it is quickly absorbed by the mucous membranes, as highlighted by the response given to addicts when cocaine is sniffed. The use of cocaine is limited, exclusively externally, as a surface analgesic, especially in the field of ENT. The maximum dose of local application must not exceed 30 mg for one use and 60 mg for 24 hours. <b>Pharmacokinetics.</b> It is easily absorbed at the level of mucous membranes. It is biotransformed intensely by hydrolysis in the blood, liver and gastrointestinal tract (when administered this way) forming benzoylecgonine (and methanol) and subsequently benzoic acid and ecgonine. It is eliminated through the kidneys. <b>Toxicodynamics.</b> Cocaine is toxic to the CNS, producing stimulating effects followed by depression, by depressing the vital centers of the bulb. Produces hyperthermia. At the level of the vegetative nervous system, it has a sympathomimetic action, the main effects are vasoconstriction and mydriasis. By repeated administration the tolerance increases, familiarity easily being set up, with a strong psychological dependence, the abstinence syndrome is insignificant in these events. The lethal dose (for unusual subjects) is smaller by mucous membrane or parenteral administration than orally. It is admitted that lethal doses for adults are 0.2 to 0.3 g s.c. and 1 g orally. <b>Treatment.</b> In acute poisoning by ingestion gastric lavage is dome with a colution of KMnO4 196 followed by the
	is done with a solution of KMnO4 1‰, followed by the administration of a purgative. Symptomatic treatment
	cardio-circulatory assistance.

Cocaine	It is presented in the form of crystals or white crystalline
hvdrochloride	powder, soluble in water (1: 0.5), alcohol (1: 3.5).
nyaroomorrae	chloroform $(1.15)$ and glycerin It has a sympathomimetic
	action by blocking programmic rountaly of poradronaling
0000	action by blocking presynaptic reuptake of norautenanne
e el	by receptors. Local anesthetic action is strong but limited
	by its toxicity; it is also vasoconstrictive locally, small
H <sub>3</sub> C-N - 0	doses causing psychomotor stimulation. It acts on the
CI- 0	central nervous system, as a stimulant in the first phase,
	and then causes depression. Produces cocainomania.
	Inside the organism, cocaine breaks down quickly, it is
	eliminated nartially through urine as econine and
	bongoig agid. Its toyigity provented its use with other
	benzoic actu. Its toxicity preventeu its use with other
	purposes apart from local anesthesia and even this use is
	limited because of fear of producing systemic effects and
	addiction.
	It is rarely used. It has the same uses as cocaine
Cocaine nitrate	hydrochloride. Due to the fact that the use of cocaine
	therapeutically is limited, presenting a number of
	disadvantages including euphoria and pronounced
	toxicity (cacoinomania): synthesis was used to obtain
H30-12 0 0 06 5	compounds with similar structure and activity Research
NO <sub>3</sub> O	has shown that most compounds in this class can fit into a
	as shown that most compounds in this class call lit life a
	general formula, which represents only part of the cocaine
	molecule responsible for local anesthesia.

Each structural component (aryl and aminoalkyl) contributes to the stability of the molecule in lipids and it can be modified to form derivatives with an increased coefficient of distribution. The substitution of the aryl radical with alkyl, alkoxy and alkylamino leads to homologous series in which the partition coefficients increase with the number of methylene groups (-CH2-). A review of these series showed that the local anesthetic activity reached the peak for the homologues C4, C5 or C6 according to the specific nature of the series under consideration. Similarly, changes in the amino alkyl group of the molecule lead to an increase of the activity and toxicity, with the increase of the number of carbon atoms. The N-alkyl branching is often accompanied by an increase of the action. Tertiary amino group may be diethylamino, piperidino or pyrrolidino, resulting in compounds that have the same degree of effectiveness. Most local anaesthetics have PKa values between 8 to 9.5. The result is that some of the compounds with higher pKa values are ionized at a rate of 100 / pH physiologically, and thus have difficulty in achieving biophase. Substances that have a lower pHa are not sufficiently ionized and are less effective, even if they reach biophase.

# **Table 2.** Local anaesthetics of synthesis and semi-synthesis [1,2,3,5,6]

Local anaesthetics and	Effect on the human body
AMINO ALCOHOLS ESTI	ERS
BENZOIC ACID DERIVATIVES (BENZOATE)	
Amylocaine (DCI), Amileina, Stovaina, Benzoate 1-Ethyl-1- methyl-2- (dimethyamino)-ethyl; 1-Dimetilamoni-2- methyl-2- benzoyl oxibutan (hydrochloride).	It is a local anesthetic as active as cocaine, but less toxic. Is also has cardiac analeptics properties; also, it does not produce arterial hypertension. It is administered internally in doses of 0.01-0.1 g per day in potions and externally, in the form of solutions 5- 20%, 4% eye drops, 1-2% ointments and 0.02-0.04 g suppositories. It can be administered parenterally (s.c.).
<i>Isobutene</i> (DCI), Benzoate 2 (isobutylamine)-2- methyl-1-propanol (hydrochloride).	White crystalline powder, easily soluble in water, alcohol and chloroform. From the structural point of view, Isobutene differs from Meprylcaine in that it has an N-isobutyl group in the place of the N-propyl group. As Meprylcaine, the action period is short. It is used in dentistry in the form of a solution of 2%.
Hexylcaine (DCI) Cyclaina, benzoate 1- (cyclohexyl- amino) -2-propanol (hydrochloride).	It is used as a local anesthetic (1-5%), with the same intensity as Cocaine and Butacaine, and for the nerve blocking anesthesia a solution of 1-2% is used, its toxicity is comparable with that produced by Procaine and Tetracaine. It is also used in spinal anesthesia with a solution of 1-2.5%.
<i>Piperocaine</i> (DCI), Metycaine, metilpiperidilpropil benzoate; 3 pipecolinopropyl benzoate; benzoate 3- (2'-methylpiperidino)	It is a crystalline substance soluble in water (1: 1.5) and alcohol (1: 4.5). Aqueous solutions are slightly acidic and stable to sterilization through autoclaving. It is a very good local anesthetic recommended as solutions of 2-4% in ophthalmology, in ENT with solutions of 2- 10%, for infiltrations are recommended solutions of 0.5-1%, and for spinal anesthesia solutions of 1,5% are used without exceeding 1.65 mg / kg.

propyl (HCl)	
CoH3-COO-CH3CH2CH2+R	
Cyclometicaine (DCI)	Cyclometicaine sulphate is in the form of a white
Surfacaine, benzoate 3-	crystalline powder, slightly soluble in water, alcohol
(2-	and chloroform.
methylpiperidino)	It is an effective local anesthetic on altered or diseased
propyl-p-	skin and on rectal mucosa. It is used topically on burns
(cichlohrxiloxi)	(sun) light skin injuries, on the rectal mucosa in the
(sulphate).	form of ointments, gels, creams or suppositories. Not
	recommended in ophthalmology.
-0-COO-CH2CH2CH2-N	
· · ·	
DERIVATIVES OF P-AM	INOBENZOIC ACID (P-AMINOBENZOATE)
<b>B</b> onzogging (DCI)	It is a local non-toxic anesthetic, readily absorbed
An eath eaine	through the altered skin and through mucous
Anestnesine,	membranes. It acts only as long as it is in contact with
Aminohonzooto othul	the diseased skin or the mucous membranes. It is used
	externally in the form of ointments and creams at
	concentrations of up to 20%, 2% oil solutions and
	suppositories. Internally it is administered as a potions
NHo	in doses of 0.3-0.6 g for ulcer pain, gastralgia.
	Procaine is the salt of a strong acid with a weak base.
	The aromatic amino group causes the aqueous solution
	to be neutral, therefore it is well tolerated by tissues.
Procaine (DCI),	The ester group may undergo hydrolytic cleavage in p-
Novocain, Allocaine,	aminobenzoic acid and diethylaminoethanol. Procaine
Sy,caine, 4-amino	hydrochloride is not effective on intact skin or mucous
benzoic 2-	membranes, but it acts promptly through infiltration.
dimethylamino acetate	Its action may be extended by concomitant
(nydrochioride).	administration with a vasoconstrictor (adrenaline) that
COO-CH2CH2-NC2H2	slows its release into the blood where it is rapidly
	inactivated by hydrolysis Procaine. It is used as a local
·HCI	anesthetic for a relatively short duration, used more in
NUL .	infiltration anesthesia, truncal, epidural and spinal
NIT2	anesthetic being weak. Administered by injection (i.v.),
	Procaine has an analgesic effect, causing vasodilatation
	and smooth muscle relaxation, depresses the heart and
	prevents certain ectopic arrhythmias, reduces the

	reflectivity of vegetation. Low doses over time, prevent aging, being a trophic sealant.
Chloroprocaine (DCI), Nesacaine, 2-Clhoro-4- aminobenzoate of diethylaminoethyl (hydrochloride). $COO-CH_3CH_2-N < C_2H_3 C_2H_3 - HCl+HCl$	Chloroprocaine differs structurally from procaine by the fact that it has an atom of chlorine in the position 2 of the aromatic nucleus which, attracting electrons, de- stabilizes the ester group. Consequently, chloroprocaine is hydrolyzed in plasma four times quicker than procaine; the local anesthetic action is more rapidly installed and is more intense than that of procaine, but the toxicity is higher.
<b>Oxybuprocaine</b> (DCI), Novesine, Butoxyprocaine, Benoxinal, 3-Butoxy-4- aminobenzoate of 2- (dietilamino)-ethyl (hydrochloride) $COO-CH_2CH_2-NC_2H_3$ +HCI +HCI +HCI	White, crystalline powder with a salty taste, very soluble in water and chloroform, soluble in alcohol, stable in air, light and heat. It appears that the 3-butoxy radical stabilizes the molecule, so it is hydrolyzed more slowly than Procaine, the degree of hydrolysis depending on the pH of the solution (pH $\approx$ 4). Oxybuprocaine is a rapidly acting surface anesthetic. By blocking reversible transmission of stimuli via sensory nerve, it is more effective than cocaine, does not cause mydriasis and accommodation disorders, being better tolerated than cocaine or tetracaine. It is indicated for surface anesthesia of the cornea, in the extraction of deep foreign bodies, in diagnostic examinations and eye surgery. It is administered as a 0.4% solution (5 drops in 5 minutes). The application of oxybuprocaine drops must be strictly limited to preparing ophthalmic surgery or diagnostic examinations. Used in small quantities, oxybuprocaine is well tolerated; sometimes causing a burning sensation, transient hyperemia, even corneal epithelial lesions. It does not cause local irritation, vasoconstriction, specific pupil dilation or sensitivity to light.
<b>Propoxycaine</b> (DCI), Blockain, 2-proxy 4-aminobenzoate of	White crystalline substance, readily soluble in water, soluble in alcohol. 2% aqueous solutions have pH 5.4. The popoxy- group apparently destabilizes the ester
2-(dimethylamino)- ethyl (chlorhydrate)	group in the same way as the chlorine in the 2-position of the chloroprocaine's structure. This is in contrast to the apparent stabilizing effect of the butoxy group
-OC <sub>3</sub> H <sub>7</sub> (n) -HCl	located in the 3-position on the aromatic ring. (Oxybuprocaine). The inductive and steric effects of a 2- alkoxy substituent which favors hydrolysis, apparently exerts greater influence than the positive mezomer

	stabilizing effect (resonance). These effects are minimal when the -alkoxy substituent is in position 3. Propoxycaine causes a more rapid onset of the local anesthetic effect and a longer duration of action, being more active than Procaine. The lipophilic nature justifies these advantages. It is administered injectable for nerve block and infiltration anesthetic, in the form of a 0.5% solutions, without the addition of a vasoconstrictor.
P-ALCOXYBENZOIC ACI	D DERIVATIVES (P-ALCOXYBENZOATES)
Paretoxycaine (DCI), Intracaine, Maxycaine, p-Ethoxybenzoate of 2-(diethylamino)-ethyl	The presence of the –alkoxy group instead of the – amino group makes Paretoxycaine produce less significant side effects than the esters of the p- aminobenzoic acid. It is used as a local anesthetic in ENT.
AMIDES	
Lidocaine (DCI), Xiline, Xylocaine,a- (Diethylamino)-2, 6- dimethylacetanilide, a- (Diethylamino)-N-(2,6 dimethylphenil)- acetamide (hydrochloride). $H_{H_{3}C} - CO-CH_{2} - N < C_{2}H_{3}$ $H_{3}C - H_{3}C - H_{3}C$	Löfgren (1940) has researched and selected Lidocaine as the main representative of the series. Based on the iodine derivative (a- diethylaminomethylindol) with low anesthetic properties, he obtained a local anesthetic with the chemical structure similar to that of lidocaine. It has the form of a white or slightly yellow crystalline powder, insoluble in water, very soluble in alcohol and oils. Lidocaine hydrochloride is a white crystalline powder, odorless, very soluble in water and alcohol. Lidocaine (xiline) is resistant to hydrolysis because, in addition to the relative stability of the amide bond, the two methyl groups at position 2 and 6 cause the steric hindrance to the attack the carbonyl group; this explains the prolonged action. Lidocaine acts on all the nerve fibers, as it contains in its molecule a hydrophilic end represented by an amino group and a hydrophobic end represented by an aromatic radical, joined by a short chain, typically ester or amide. In tissues due to the slightly alkaline pH and buffer system, the liposoluble base is released, which is able to penetrate through the membrane of nerve fibers, where the amide undergoes a quartemizing effect. Lidocaine is used in all types of local anesthesia it is three times

	more active than procaine and has longer action. As antiarrhythmic, it is used in ventricular arrhythmia in acute myocardial infarction, cardiac surgery or therapy with digitalis, being administered i.v., typically under EKG control, from the initial dose of 100 mg, to the reach of the effective plasma concentration, continuing with 1.5-3 mg / minute.
<b>Bupivacaine</b> (DCI), Sensorcaine, Marcaine, Carbostesia, (±)-N-(2,6- Dimethylphenyl)-1- butyl-2- piperidincarboxamide (hydrochloride).	Bupivacaine is a local analgesic characterized by a long duration of action, and a relatively high installation latency. The pharmacodynamic properties are similar to lidocaine. The effect occurs slowly, in 15 minutes, and lasts 4-8 hours, due to its marked liposolubility and to its ability to bind to membrane proteins. Bupivacaine is used in local or regional anesthesia for surgery (truncal, plexicală, caudal, epidural anesthesia) and is administered parenterally; doses should not exceed 2.5 mg / kg body weight.
Mepivacaine (DCI), Carbocaine, Mepivastein, Opticain, Scandonest, (±) N-(2,6- Dimethylphenyl)-1- methyl-2-piperidine- carboxamide (hydrochloride); 1- Methyl-2,6'- pipelocoxylidide (hydrochloride).	As with the other anilides and lidocaine, mepivacaine is highly resistant to hydrolysis, and its solutions are subjected to sterilization in autoclave, without decomposition. Mepivacaine is used as racemic because it was established that the two optical isomers have the same activity and toxicity; they are comparable to those of lidocaine. Its duration of action is considerably longer than that of lidocaine, even in the absence of a vasoconstrictor. It is recommended especially when epinephrine and its counterparts are contraindicated. Mepivacaine is a local anesthetic of synthesis whose action is due to the decrease in membrane permeability of sodium ions. It is administered parenterally 1 to 2 ml of a 3% solution.
<b>Prilocaine</b> (DCI), Citanest, N-(2- Methylphenyl)-2- (propilamina)- propanamide (chlorhydrate); 2- (Propilamino)-o- propionotoluidide	White crystalline powder, odorless, bitter, slightly soluble in water and alcohol. Prilocaine has stability, effectiveness, toxicity and duration of action similar to that of other anilides. Its duration of action is between that of lidocaine and that of mepivacaine. Side effects are similar to those produced by other anilides, noting that prilocaine produces methemoglobinemia. This is due to the fact that as a result of its metabolism is obtained a-toluidine- (2-methylaniline), which is

European Journal of Natural Sciences and Medicine

(hydrochloride)	broken down into toxic products
CH3	(phenylhydroxylamine nitrobenzene) that cause
NH-CO-CH-NH-CH <sub>2</sub> CH <sub>2</sub> CH <sub>3</sub>	methemoglobinemia. Prilocaine hydrochloride
HGL HCL	solutions are used without addition of vasoconstrictors.
ETHERS	
Pramoxin (DCI),	White crystalline powder with aromatic smell, slightly
Pramocaine,Tronothan	soluble in water and alcohol, the pH of THE 1% solution
e,4-[3-(p-	is 4.5.
Butoxyphenoxy)-	It is a topical local anesthetic, with a low index of
propyl]-morpholine	sensitivity, with few side effects. It is used for relieving
(hydrochloride)	pain and itching caused by insect stings lesions,
· 0CH2CH2CH2-N	superficial injuries and hemorrhoids.
$OC_4H_2(n)$	
OTHER STRUCTURES	
Foncaina (DCI)	White crystalline powder, odorless, bitter-tasting,
Holocoine N N'-bis-(A-	stable in air, easily soluble in water (1:50), easily
Ethoyunhenul)	soluble in alcohol. The aqueous solutions may be
acetamidine	sterilized by boiling, but are unstable to the action of
monohydrate	alkaline hydroxides. The local anesthetic properties of
(hydrochloride)	fencaine were highlighted in 1897, preceding the
(ilyuroemoride)	discovery of izogamine.
$NH-C=N-OC_2H_5$	Fencaine hydrochloride is a slightly irritating, causing a
ĊH <sub>3</sub>	slight discomfort before the onset of anesthesia. It is
• HCl	more toxic than cocaine, and therefore cannot be used
OC2H4	for parenteral administration; it is used as a $1\%$
	ophthalmic solution 1 to 2% ointment.

The spectrums obtained for the analyzed substances (Cocaine, Benzocaine, Procaine, Lidocaine, Bupivacaine, Prilocaine) are presented in the following figures. *IR* spectrums of the analyzed compounds are presented by describing the transmittance depending on the wavelength.



### Fig. 1 IR spectrum of cocaine







Fig. 3 IR spectrum of procaine







Fig. 5 IR spectrum of bupivacaine





# Conclusions

Local anaesthetics may have 3 types of systemic effects, namely: central-nervous, vegetative and cardiovascular. Being absorbed at the site of applications, they cause central excitement phenomena, followed by depression. These effects may lead to serious toxic symptoms. The slowing of absorption by associating vasoconstrictors allows to prolong local action and attenuates resorption effects:

- a) at the level of the central nervous system, excitation and depression events may take place, sometimes a state of confusions (similar to intoxication), after which may follow coma and finally the paralysis of respiratory centers.
- b) at the level of the vegetative nervous system they produce a vasodilator, hypotensive effect, also having a certain antispasmodic action at the level of the smooth muscles.
- c) at the level of the cardiovascular system they cause myocardial depression by inhibiting excitability. In overdose, myocardial depression may be severe. But this effect is therapeutically exploited in cardiac arrhythmia.

Local synthesis and semi-synthesis local anaesthetics have multiple uses in dentistry, ENT, surgery and obstetrics, representing very useful "weapons" in the fight against pain.

The obtained specters may bring a contribution to the setup of an atlas of specters to be used in the control of medicines.

### Acknowledgement

This paper is supported by the Sectoral Operational Programme Human Resources Development (SOP HRD), financed from the European Social Fund and by the Romanian Government under the contract number POSDRU/187/1.5/S/156040/

### References

- [1] Aurelia Nicoleta Cristea "Tratat de Farmacologie Ediția I" Medical Publishing House Bucharest 2005
- [2] Cristea L., Noul ghid de Anestezie Terapie Intensivă. Medical Publishing House Bucharest, 1992
- [3] Elena Hațieganu; Camelia Stecoza "Chimie Terapeutică Vol. 2" Editura Medicală Bucharest 2010
- [4] Sîrbu R., Negreanu-Pîrjol T.: Metode fizico-chimice de analiză a substanţelor farmaceutice, Editura Matrix Rom, Bucharest, 2004.
- [5] H. Gall, R. Kaufmann, C.M. Kalveram, Adverse reactions to anaesthetics: analysis of 197 cases, J. Alergy Clin. Immunol. 4 (1996)
- [6] Becker, DE; Reed, KL: Essentials of Local Anesthetic Pharmacology. Anesth Prog 53:98-109 2006