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0-Surgery

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Lec: 4

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## Vasoconstrictor agents

Vasoconstrictors are drugs or chemical agents that constrict blood vessels thereby control tissue perfusion, They are added to local anesthetic solutions to oppose the vasodilatation actions of local anesthetics. All inject able local anesthetics posses some degree of vasodilating activity. The degree of vasodilatation varies from one type to another and may vary according to the site of injection and individual patient response.

After local anesthetic injection into tissue, blood vessel dilate in the area, resulting in an increased blood flow to the site. This increase in perfusion leads to the following reactions:

- 1 increased rate of absorption of the local anesthetic into the cardiovascular system ,which in turn removes it from the injection site.
- 2- increased plasma level of the local anesthetic , with an increased risk of local anesthetic toxicity .
- 3 decreased duration of action and decreased depth of anesthesia because it diffuses away from the injection site more rapidly.
- 4 increased bleeding at the site of injection due to increased perfusion.

# The advantages of additions of vasoconstrictors to local anesthetic solution are:-

- A by vasoconstricting blood vessel , vasoconstrictors decrease blood flow ( perfusion ) to the site of the injection .
- B absorption of the local anesthetics into the cardiovascular system is slowed ,resulting in lower anesthetic blood levels .
- C the lower local anesthetic blood levels decrease the risk of local anesthetic toxicity .
- D higher volumes of the local anesthetic agent remain in and around the nerve for longer periods thereby increasing the duration of action of local anesthetics .

E - vasoconstrictor decrease bleeding in the site of injection and are useful when increased bleeding is anticipated (ex: during a surgical procedure).

### Classification of vasoconstrictor

These drugs can be classified on the basis of mode of action , into the following three categories :

- 1- Direct acting drugs : these drugs stimulate or exert their action directly on the adrenergic receptors , for example epinephrine , nor epinephrine , levonordefrine .
- 2- indirect acting drugs: These drugs act by releasing nor epinephrine from the adrenergic nerve terminals, for example tyramine, amphetamine.
- 3 Mixed acting drugs: These drugs have both direct and indirect actions, for example ephedrine.

All the vasoconstrictors used in conjunction with local anesthetic are direct acting agents .

#### Dilution of vasoconstrictors:

The dilution of vasoconstrictor is commonly referred to as ratio (1 to 1000, and is written as 1:1000). This 1:1000 mean that there is I gm (1000 mg) of the drug contained in 1000ml of solution.

## Type of the vasoconstrictors :-

## A – Epinephrine (adrenaline )

Epinephrine remains the most effective and the most commonly used vasoconstrictor in medicine and dentistry.

**Sources**:- it is secreted primarily by adrenal medulla. It is available as synthetic and is also obtained from adrenal medulla of animals.

Mode of action : - it acts directly on both alpha and beta adrenergic receptors .

## Systemic action:-

1 - cardiovascular system

It causes increased systolic and diastolic blood pressure ,increased heart rate and strength of contraction ,increased stroke volume and cardiac output ,and it causes increased myocardial Oxygen consumption On blood vessel it cause vasoconstriction and so frequently used a lone as a vasoconstrictor for hemostasis during surgical procedures .

2 - respiratory system

Adrenaline is a potent dilator of the smooth muscle of the bronchiole , so it is the drug of choice for management of acute asthma .

3 - central nervous system

In the usual therapeutic dosage adrenaline is not a potent CNS stimulant, CNS stimulation occurs when an excessive dosage is given .

#### Availability in dentistry

Adrenaline is the most potent and widely used vasoconstrictor in dentistry . it is available in 1 :50 000 , 1 : 80 000 ,1 : 100 000 ,  $1:200\ 000\ dilution$ 

#### Maximum dosage:

This drug is potent, and can produce undesirable results

A – if used in large volumes

B – if inadvertently injected intravascularly .

So these drugs should be used with consideration to their benefits and risks.

The maximum doses are recommended by New York heart association, and as suggested by Bennette (1983) and Malamed (1977) are as follows:

A - For normal healthy adult patient the maximum recommended dose is (0.2 mg) per appointment that means

10 ml of a 1: 50 000 dilution (5 cartridge)

16 ml of a 1 : 80000 dilution (8 cartridge)

20 ml of a 1:100000 dilution (10 cartridge)

40 ml of a 1:200 000 dilution (20 cartridge)

# B - For patient with clinically significant cardiovascular disease The max dose is 0.04 mg per appointment, that means

2 ml of 1:50 000 dilution (1 cartridge)

3.2 ml of a 1:80000 dilution (1.6 cartridge)

 $4 \text{ ml of a } 1:100\ 000\ \text{dilution}$  (  $2\ \text{cartridge}$  )

8 ml of a 1 : 200 000 dilution ( 4 cartridge ) .

## B- Nor epinephrine (nor adrenaline)

Source: either synthetic or obtained from adrenal medulla of animals.

**Mode of action**: it act almost exclusively on alpha receptors. it also stimulate beta receptors in the heart, it has one fourth of adrenaline potency.

#### Systemic action

1 – on cardiovascular system

It causes increased systolic blood pressure, decreased heart rate, slightly decrease in cardiac output ,increased stroke volume. On the blood vessel it cause vasoconstriction.

2 – on respiratory system

It dose not relax the smooth muscle as do adrenaline and it is not clinically effective in the management of bronchospasm.

3 - on CNS

In the usual therapeutic dose nor epinephrine is not potent CNS stimulant.

#### Availability in dentistry:

Nor epinephrine is used with local anesthesia as vasoconstrictor in a 1:  $300\,000\,\text{dilution}$  .

Maximum recommended dose,: -

for normal healthy adult  $\,$  patient the maximum recommended dose is 0 . 34 mg per appointment , that mean 10 ml of a 1:  $\,300\,000$ 

For patient with clinically significant cardiovascular disease

0. 14 mg per appointment that means 4 ml of a  $1:300\ 000$  .

## C - Levonordifrine

It is synthetic substance, it act through direct alpha receptor stimulation with some beta activity , it produces less cardiac and CNS stimulation than epinephrine dose it is mainly used  $% \left( 1,0\right) =0$  with mepivacaine in a  $1:20\;000$ 

the maximum dose for all patient should be 1 mg per appointment that mean 20 ml of a 1:20000 dilution (11 cartridge).

## D - Felypressine

Source : it is a synthetic analogue of the anti diuretic hormone (vasopressin ) . it is a non sympathomimetics drug and is categorized as vasoconstrictor.

Mode of action: - it act as direct stimulant of vascular smooth muscle.

Systemic action :-

Heart: no direct effect.

Blood vessel: in high doses Felypressine - induced constriction of the cutaneous blood vessel.

CNS: no effect.

Uterus: it has oxytocic action so it is contraindicated for pregnant patients.

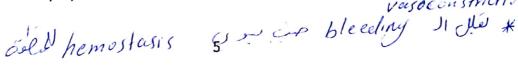
# Availability in dentistry

It is used in dentistry in a dilution of 0.03 IU / ml with 3 % prilocaine. Maximum dose is 0.27 IU that's mean 9 ml of 0.03 IU / ml.

#### Selection of a vasoconstrictor

The selection of an appropriate vasoconstrictor is based on following factors:

- 1 Length of the surgical or dental procedure .
- 2 Requirement for hemostasis during the surgical procedure.
- 3 Requirement for postoperative pain control .
- 4 Medical or physical status of the patient and if used any medications



# 1 – Length of the surgical procedure

The duration of pulpal and hard tissue anesthesia with 2% lidocaine only lasts for 10 min , the addition of adrenaline of  $1:50\ 000$  or  $1:100\ 000$  dilution prolong the duration to 60 min ( Newcomb and Wait 1972 ).

Hence for any surgical treatment requiring 40 -50 min it is difficult to achieve pulpal anesthesia without using a vasoconstrictor .

## 2 – Requirement of hemostasis

Some of the vasoconstrictors are effective in minimizing the blood loss during the surgical procedure as adrenaline , others as Felypressine which is of minimal value in achieving hemostasis .

## 3 - Requirement of postoperative pain control

Profound pain control of adequate duration by a local anesthetic agent with a vasoconstrictor is used .Plain local anesthetic agent produce pulpal anesthesia of shorter duration than that of local anesthetic agent with vasoconstrictor and is likely to produce stress response .

4 — Medical or physical status of the patient and medication used The benefit and risks of including a vasoconstrictor in a local anesthetic solution in patients who are medically compromised ., must be weighed against benefits and risks of using plain anesthesia .

For the following group of patients the use of local anesthetic agents with vasoconstrictor is contraindicated :

- A Patients with significant cardiovascular disease such as ischemic heart disease, hypertension, cerebral strokes.
- B Patients with certain uncontrolled non cardiovascular disease such as thyrotoxicosis or hyperthyroid states.
- ${\sf C}$  Patients receiving monoamine oxidase inhibitors , tricyclic antidepressant .

## Reducing agent

Vasoconstrictors in local anesthetic solution are unstable and maybe oxidize especially on prolonged exposure to sunlight and this will lead to brown discoloration of the solution , to overcome this problem a small quantity of antioxidant as sodium bisulphate is added to the cartridges ,this substance reacts with Oxygen before oxygen can destroy the vasoconstrictor ( adrenaline or nor adrenaline ) so it protects their stability.

#### **Preservative**

The sterility of local anesthetic solution is maintained by the inclusion of small amount of preservative . A number of chemicals are used as general preservatives, these are added to increase the shelf –life, and include:

- 1 Methylparaben which is bacteristatic and fungistatic agent
- 2 Thymol which is antiseptic, fungistatic.

#### **Fungicide**

In the past some solutions tended to become cloudy due to the proliferation of minute fungi, now a small quantity of thymole is added to serve as a fungicide and prevent this occurrence.

## Sodium chloride (Nacl) or Ringers solution

The anesthetic agent and the additives mentioned above are dissolved in modified ringers solution. It is added to content of the cartridge to make the solution isotonic with the tissue of the body This isotonic vehicle minimizes the discomfort during injection of local anesthesia.

#### Distilled water

It is used as a diluents to provide the volume of solution in the dental cartridge.