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Comparative Evaluation of Hemodynamic Stability and Recovery During Conscious Sedation by Dexmedetomidine-Fentanyl Versus Ketamine- Fentanyl in procedures outside the operating room.

Thesis

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List of Contents

Title	Page No.
Introduction	1
Aim of the Work	3
Review of Literature	
Conscious Sedation	4
Dexmedetomidine	Error! Bookmark not defined.
Ketamine	Error! Bookmark not defined.
• Fentanyl	Error! Bookmark not defined.
Patients and Methods	31
Results	38
Discussion	43
Summary	49
Conclusion	52
References	53
Arabic Summary	



INTRODUCTION

Cognizant mild is a process of giving lacking of pain, mild and portion while guaranteeing fast recuperation without reactions. Cognizant mild is controlled with the double objectives of quickly and securely building up palatable procedural condition for the exhibition of helpful or demonstrative methods while guaranteeing fast, unsurprising recuperation with insignificant post-usable spin-offs (Jalowiecki et al., 2005).

Dexmedetomidine is particular $\alpha 2$ fact agonist that has narcotic, sympatholytic, amnestic and torment relieving impacts. It has been set in various manual preliminaries as valuable and safe substance. Giving a great lacking of torment, cognizant mild in sick who appear to be snoozing, anyway can undoubtedly be stir, notwithstanding no respiratory misery, make component one of the generally utilized drug in mild (Panzar and Moitra, 2011).

These properties of component render it appropriate for mild and lacking of torment during the entire perioperative period. Its applications as a premedication, as a soporific aide for general and territorial mild and as a postoperative soothing



and torment relieving are like those of the structures (Cheung et al., 2007).

AIM OF THE WORK

The point of this examination is to research whether Component with Agent mix is a successful elective methodology to Drug with Agent as respect mild, partials soundness and recuperation for sick experiencing techniques outside the working room, for example, obtrusive radiology, bronchoscopy, gastroenterology and mental dealing (ECT).

Chapter 1

Conscious Anesthesia

Cognizant mild is characterized as a therapeutically controlled condition of discouraged awareness wherein sick hold their defensive reflexes, keep up their aviation route freely, and react to physical and verbal incitement. Cognizant mild is moderate in the range of mild, which extents from portion and lacking of torment to general mild. The key qualities of cognizant mild are that it is of fast beginning, is titratable to an individual sick's ideal degree of mild, and is related with huge wretchedness of awareness, help of nervousness, and lacking of torment. Retrograde amnesia of any agony related with techniques is an extra advantage of cognizant mild. The impacts of the prescriptions most regularly utilized for cognizant mild are reversible with pharmacological foes (Clark and Tri, 2000).

Meaning of general mild, level of mild:

Insignificant Mild (Portion): is a medication actuated state during which sick react regularly to verbal directions. Albeit subjective capacity and physical coordination might be hindered, aviation route reflexes, and ventilator and cardiovascular capacities are unaffected. (Cote and Wilson, 2016).

Moderate Mild/Agents ("Conscious Mild"): is a medication incited wretchedness of cognizance during which sick react intentionally to verbal directions, either alone or joined by light material incitement. No intercessions are required to keep up a patent aviation route, and unconstrained ventilation is satisfactory. Cardiovascular capacity is generally kept up (Coté CJ, 2016).

Profound Mild/Agents:- is a medication incited wretchedness of cognizance during which sick can't be effectively excited however react intentionally following rehashed or difficult incitement. The capacity to autonomously keep up ventilatory capacity might be hindered. Sick may require help with keeping up a patent aviation route, and unconstrained ventilation might be lacking. Cardiovascular capacity is generally kept up (Coté CJ, 2016).

General Mild:- is a medication actuated loss of cognizance during which sick are not arousable, even by excruciating incitement. The capacity to autonomously keep up ventilatory capacity is frequently impeded. Sick regularly

require help with keeping up a patent aviation route, and positive weight ventilation might be required in light of discouraged unconstrained ventilation or medication incited sadness of neuromuscular capacity. Cardiovascular capacity might be weakened (Cox B, Durieux ME, 2003).

Part 2

COMPONENT

(α2-fact agonist)

T

he $\alpha 2$ receptors are situated on veins, thoughtful terminals, focal sensory system, and their enactment prompts mild, a decrease of tonic degrees of thoughtful surge, and an increase of cardiovascular vagal movement (Thomas et al., 2000).

Pharmacology of component

Component is an imidazoline subordinate. Its synthetic structure is 4-(S)- [1-(2,3-dimethylphenyl)ethyl]-1-H-imidazole mono hydrochloride. Component is tolerably lipophilic (Figure 1) (Kuusela et al., 2001).

Pharmacokinetics:-

o Absorption:

In spite of the fact that component is enrolled for IV use, numerous courses of organization have been investigated. After oral organization, a broad first-pass impact is watched, with a bioavailability of 16%. Component is all around ingested through the intranasal and buccal mucosae, an element that could be of advantage when utilizing component in unco-processyoungsters or geriatric sick (Li et al. 2016).

o Distribution:

Component is a very protein-bound medication. In plasma, 94% of component is bound to egg whites and $\alpha 1$ -glycoprotein. In pre-manual creature ponders, it was discovered that component promptly crosses the blood–cerebrum and placenta obstructions Using non-compartmental examination, a circulation half-existence of around 6 min was found in sound volunteers . After long haul imbuement in ICU sick with hypoalbuminemia, an expanded volume of circulation at consistent state was watched (Välitalo PA et al ,2013) .

o Metabolism:

Component is disposed of essentially through biotransformation by the liver. A hepatic extraction proportion of 0.7 was found . Under 1% is discharged unaltered with metabolites being discharged renally (95%) and fecally (4%) .

Direct N-glucuronidation by uridine 5'-diphospho-glucuronosyltransferase (UGT2B10, UGT1A4) represents about 34% of component digestion. Moreover, hydroxylation intervened by cytochrome P450 (CYP) proteins (for the most part CYP2A6) was exhibited in human liver microsomes (Lee et al ,2012)

o Elimination:

The hepatic extraction proportion of Component has been recently assessed at 70%, so changes in local hepatic blood stream may affect Component pharmacokinetics, however the impact is little. Past research has demonstrated that 19% decline in heart yield brought about an expected 12% diminishing in freedom. Changes in renal blood stream would not be required to influence pharmacokinetics Additionally, the pharmacokinetics in subjects with ordinary renal capacity, gentle, moderate and extreme renal disability, as characterized by creatinine freedom, did not vary (Venn et al., 2001).

For component, drawn out just as abbreviated end halflives have been accounted for sick with hypoalbuminemia. Leeway be that as it may, is just imperceptibly influenced by hypoalbuminemia (Zhang et al ,2015).

Pharmacodynamics:-

1-Agents

They drag out the length of lacking of torment by obstructing the hyperpolarization-enacted cation current (Ih current). This is the most well-characterized component of α 2-Adrenoreceptor agonists. After an activity potential (AP) has happened, the nerve should repolarize to have the option to create new APs. The early repolarization stage will result in a hyperpolarized express that will make the age of new APs for all intents and purposes unimaginable, and the nerve is, during this period, made a decision to be unmanageable to incitement. In this manner, hindering the Ih current will bring about delayed hyperpolarization of the nerve, which thus will bring about a torment relieving activity. Hindering the Ih current may likewise can possibly deliver a particular tangible impact as this impact seems, by all accounts, to be progressively articulated in C filaments (torment) than in An alpha strands (engine). Component has progressively articulated impact on hindrance of nerve fiber

activity potential when contrasted with clonidine (Lönnqvist PA. 2012).

Midway, $\alpha 2$ -AR agonists cause lacking of torment and mild by hindrance of substance P discharge in the nociceptive pathway at the degree of the dorsal root neuron and by enactment of $\alpha 2$ -AR in the locus coeruleus. Concealment of action in the sliding noradrenergic pathway, which adjusts nociceptive neurotransmission, ends engendering of torment sign prompting lacking of torment. (Grewal A . 2011)

Another conspicuous physiologic activity credited to $\alpha 2$ facts is their decrease of calcium conductance into cells, in this way hindering synapse discharge. This impact includes direct guideline of calcium passage through N-type voltage-gated calcium channels and is autonomous of cAMP and protein phosphorylation. It is intervened by G0 proteins. These 2 components speak to 2 altogether different methods for affecting lacking of torment: in the main, the nerve is kept from consistently terminating, and in the second, it can't proliferate its sign to its neighbor (Birnbaumer et al., 1990).

2- Mild

The calming activity of Component is intervened through its activity on focal α 2-facts in the locus ceruleus and receptors in the dorsal horn of the spinal string are associated with these impacts (Judith et al., 2000).

Component has been exhibited to be a powerful soothing in basic consideration settings. Early investigations contrasting premedication among Component and midazolam noticed no distinction in the nature of portion (Ramsay, 2000).

There is a portion reaction relationship that connects expanding Component plasma levels with the more prominent degrees of mild. Component has been reported as being utilized manually in portions up to 2.5 μ g/kg/hr; since the bundle addition shows a furthest point of confinement of implantation of 0.7 μ g/kg/hr, the referenced information propose that Component can be promptly titrated in its manually important range (Jaakola et al., 1994).

In spite of the fact that the component by which Component weakens excitement to create mild isn't exactly known, one perceived site of activity is the locus coeruleus (LC), a dorsolateral pontine core of roughly 30,000 neurons. So named (interpreted from Latin, the locus coeruleus is the "blue spot") due to a high centralization of the blue-recoloring