

A Clinical Assessment of the Efficacy and Hemodynamic Stability of Propofol- Ketamine and Propofol-Fentanyl in Patients Undergoing Dilatation and Curettage: A Comparative Study

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Abstract

Aim: The aim of the present study was to evaluate and compare the efficacy and hemodynamic stability of ketamine / fentanyl along with propofol and also assess recovery profile, post-operative analgesia and complications in patients undergoing Dilatation and Curettage.

Methods: The present study was carried out in the Department of Anesthesia Sri Krishna Medical College & Hospital, Muzaffarpur, Bihar, India for one year, patients written informed consent, the study was conducted in 100 patients, aged 18-35 years of ASA Grade 1&2 scheduled for Dilatation and Curettage procedures lasting up to 30 minutes.

Results: Pre-induction heart rate, systolic blood pressure, diastolic blood pressure, mean arterial pressure, spo₂ were comparable in both the groups with a statistically no significant difference between them (p<0.05).

Conclusion: It may be concluded from our present study that propofol with ketamine as an adjuvant in the dose of 0.75 mg/kg compared to propofol with fentanyl as adjuvant in the dose of 2 mcg/kg provides deep sedation. But propofol-ketamine group is more efficacious and provides better peri-operative hemodynamic stability during anesthesia as compared to propofol-fentanyl group. Also the propofol-ketamine combination produces good analgesia with less requirement of rescue drug in post-operative period with fewer peri-operative complications than propofol-fentanyl combination. We have not encountered any psychotomimetic effects of injection Ketamine in the 0.75mg/kg dose. Hence propofol-ketamine combination is a better choice especially when hemodynamic stability is of great importance in patients undergoing dilatation and curettage.

Keywords: Propofol, Ketamine, Fentanyl, Sedation, Hemodynamic stability, Dilatation, Curettage

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Introduction

Day care gynecological procedures require the use of anesthetic agents which ensure rapid induction and recovery. [1] Total intravenous anesthesia (TIVA) is a combination of hypnotic agents, analgesic drugs and may be muscle relaxants, excluding simultaneous administration of any inhaled drugs. Therefore, it can be an effective alternative to inhalational anesthesia and for ambulatory surgery when the speed and completeness of recovery are important. Drugs used for TIVA should have quick onset, smooth induction, easy maintenance, quick recovery and

minimal side effects. [2] Ideal drug for Sed analgesia should have rapid onset and fast recovery time. However, there is still no consensus for best Sed analgesia management for short-term procedures.1 Intra-venous-based anesthesia techniques are widely used for the patient who must sleep during the procedure. [3] Although propofol is the gold standard drug in day care procedures, it has its own side effects like apnea, cardio-vascular instability, pain on injection. [1] Ketamine is an agent that provides sedation, analgesia and amnesia, and it might be an appropriate option for short-

lasting procedures. However, it has cardio-vascular side effects and an induction of transitory psychotic episodes, together with delayed recovery and secretion increment. [4] Ketofol (propofol-ketamine admixture) is a combination of ketamine and propofol that is an agent of choice for various procedures. [2] The safety and efficacy of ketofol as a sedo-analgesic agent depend on the dose and the ratio of the admixture. [5]

Combining propofol with other drugs such as opioids or ketamine is recommended for improving the quality of sedation, analgesia and minimizing the potential adverse effects with maintenance of a stable cardiovascular and respiratory status in the perioperative period.² Ketamine, a NMDA receptor agonist, in subanesthetic doses with propofol has gained attention in total intravenous anesthesia because of its powerful analgesic action without causing myocardial and respiratory depression. Ketamine also causes some degree of sympathetic stimulation, which tends to counter balance the cardiovascular effects of propofol. [6] Fentanyl on the other hand is synthetic opioid analgesic which has rapid onset and short duration of action and has been used in combination with propofol satisfactorily.

Ketofol (propofol-ketamine admixture) is a combination of ketamine and propofol that is an agent of choice for various procedures. [7] The safety and efficacy of ketofol as a sedoanalgesic agent depend on the dose and the ratio of the admixture. [8] The ratios of 1:2, 1:3 and 1:4[sub dissociative dose] ratios were very effective for the day case procedure. [9] Dilation and curettage (D C), a brief and painful procedure, is performed for the diagnosis and treatment of endometrial and intrauterine disorders. The procedure is one of the most frequently performed gynecological surgical procedures. It causes considerable pain during cervical dilation and tissue extraction. [10]

The aim of the present study was to evaluate and compare the efficacy and hemodynamic stability of ketamine / fentanyl along with propofol and also assess recovery profile, post-operative analgesia and complications in patients undergoing Dilatation and Curettage.

Materials and Methods

The present study was carried out in the Department of Anesthesia, Sri Krishna Medical College & Hospital, Muzaffarpur, Bihar, India for one year, and patients written informed consent, the study was conducted in 100 patients, aged 18-35 years of ASA Grade 1&2 scheduled for Dilatation and Curettage

procedures lasting up to 30 minutes. The patients were randomly allocated using sealed envelope method in 2 groups (50 of each). Patients with respiratory, cardiovascular, neurological, liver diseases, patients on narcotic therapy were excluded from study. Pre-anesthetic evaluation including history and a thorough general and systemic examination and all relevant investigations were done for all the patients.

Patients were kept fasting for at-least 8 hours prior to anesthesia. Pre- operative baseline heart rate, systolic blood pressure, diastolic blood pressure, mean blood pressure, spo2 was recorded. A peripheral iv line was established. All patients were pre-medicated with Inj. Glycopyrrolate 0.004 mg/kg and Inj. Midazolam 0.02 mg/kg. Group A received Inj. Ketamine 0.75 mg/kg body weight IV slowly over 2 minutes, after 5 minutes Inj. Propofol given at rate of 1ml/3 seconds till loss of consciousness. Group B received Inj. Fentanyl 2 mcg/kg body weight as slow IV injection, after 5 minutes Inj. Propofol given at rate of 1ml/3 seconds till loss of consciousness.

Intra-operatively heart rate, blood pressure, oxygen saturation and Ramsay sedation score was recorded at different time intervals of 5, 10, 15, 20, 25, 30 minutes following induction of anesthesia in both groups. Throughout the procedure patients were allowed to breathe spontaneously on room air and oxygen supplementation was given to some patients during apnea.

Top-up dose of Inj. Propofol 0.5 mg/kg was given when patient became light during anesthesia as indicated by rise in heart rate, blood pressure or any other movement to surgical stimulus. Total dose of propofol required for the patients was noted.

Post-operatively all vital parameters like heart rate, blood pressure, oxygen saturation and visual analogue score was recorded every 30 minutes for first 2 hours then every 2 hours till 12 hours. Any complication like nausea, vomiting, delirium, sedation, pain was noted.

Statistical Analysis

Continuous variables like age, heart rate, systolic blood pressure, diastolic blood pressure, Ramsay sedation score, spo2, total dose of propofol etc were presented as Mean \pm SD. Continuous variables were compared at different time intervals between ketamine and fentanyl groups by performing unpaired t-test, $p < 0.05$ was considered as statistically significant.

Results

Table 1: Comparison of Heart Rate In Both Groups

Heartrate	Baseline	1 min	5 min	10min	15min	20min	25min	30min
Group A Mean± SD	83.57±3.37	82.22±4.94	86.64±5.3	83.32±6.34	82.0±5.55	81.49±6.03	80.24±6.2	80.8±6.2
Group B Mean± SD	86.82±3.87	88.06±6.06	83.67±6.00	78.22±5.25	78.34±4.86	76.45± 6.63	80.65±6.16	79.55±6.31
P-value	0.01	0.0001	0.0007	0.0003	0.0007	0.03	0.763	0.482

Table 2: Comparison of Systolic Blood Pressure (SBP) In Both Groups

SBP	Baseline	1 min	5 min	10min	15min	20min	25min	30min
Group A Mean ± SD	123.64±5.48	119.14±12.17	120.60±11.30	119.14±9.90	116.35±10.97	114.10±11.87	114.78±10.56	115.07±10.23
Group B Mean ± SD	123.35±4.52	119.85±9.67	117.21±8.35	116.57±9.19	111±7.44	108.42±8.21	109.85±8.68	107±18.77
P-value	0.823	0.803	0.191	0.301	0.030	0.04	0.05	0.001

Table 3: Comparison of Diastolic Blood Pressure(DBP) In Both Groups

DBP	Baseline	1 min	5 min	10min	15 min	20 min	25 min	30 min
Group A Mean ± SD	72.47±4.91	69.65±7.04	69.38±2.95	67.42±5.56	67.22±5.34	67.35±5.97	67.47±4.39	69.25±3.78
Group B Mean ± SD	73.37±4.51	69.77±4.88	69.47±3.51	64.75±3.59	64.45±3.59	65.58±4.71	67.37±4.39	25
P-value	0.370	0.858	0.802	0.013	0.017	0.060	0.033	0.926

Table 4: Comparison of Ramsay Sedation Score in Both Groups

RSS	Baseline	1 min	5 min	10 min	15 min	20 min	25 min	30 min
Group A Mean ± SD	1.92±0.27	1.92±0.27	4.40±0.57	4.82±0.56	5.46±0.57	5.86±0.35	5.96±0.19	5.95±0.20
Group B Mean ± SD	1.94±0.23	1.94±0.23	3.94±0.74	4.06±0.72	5.10±0.57	5.72±0.49	5.9±0.30	5.97±0.14
P-value	0.70	0.70	0.009	<0.001	<0.001	0.20	0.35	0.65

Table 5: Comparison of VAS in Both Groups

VAS	30 min	1 hour	1 Hour 30min	2hour	4 hour	6 hour	8 hour	10 hour	12 hour
Group A Mean± SD	2.87±1.22	2.33±1.12	1.57±0.9	1.33±0.76	1.17±0.53	1.1±0.4	1±0	1±0	1±0
Group B Mean ± SD	4.37±1.54	3.27±1.34	2.8±1.13	2.3±0.84	1.63±0.85	1.23±0.5	1±0	1±0	1±0
P-value	0.001	0.004	<0.001	<0.001	0.013	0.262	-	-	-

Pre-induction heart rate, systolic blood pressure, diastolic blood pressure, Ramsay sedation score, VAS were comparable in both the groups with a statistically no significant difference between them (p<0.05).

Discussion

Day care gynecological procedures require the use of anesthetic agents which ensure rapid induction and recovery. [11] Total intravenous anesthesia (TIVA) is a combination of hypnotic agents,

analgesic drugs and may be muscle relaxants, excluding simultaneous administration of any inhaled drugs. Therefore, it can be an effective alternative to inhalational anesthesia and for ambulatory surgery when the speed and completeness of recovery are important. Drugs used for TIVA should have quick onset, smooth induction, easy maintenance, quick recovery and minimal side effects. [2] Ideal drug for sedo-analgesia should have rapid onset and fast recovery time. However, there is still no consensus for best Sed analgesic management for short-term procedures. [10]

Pre-induction heart rate, systolic blood pressure, diastolic blood pressure, Ramsay sedation score, VAS were comparable in both the groups with a statistically no significant difference between them ($p < 0.05$). The minimal fall in SBP, DBP and MBP in group PK as compared to group PF is due to the sympathomimetic activity of ketamine which counteracts with the cardiovascular depressant action of propofol, thus maintaining a stable hemodynamic profile as compared to fentanyl. These findings are correlated well with the study done by Pawar et al [12], Phillips et al [13] and Khutia et al. [14] The opposing effect of ketamine and propofol on arterial pressure tended to cancel each other out resulting in improved cardiovascular stability. The propofol ketamine group experienced a smaller percentage decrease in SBP, which is comparable with other studies. [6,7,15]

The ketamine-propofol combination is thought to act by counteracting the cardiovascular side-effects of each other, preserving the sedative efficacy. Also the amount of propofol needed to achieve a deep sedation level was much lower in case of PK group than PF group, which contributed to the lower incidence of hypotension and apnea. Also, no patients in present study required airway manipulation. The post operative complications were also higher in group PF than in group PK. There were no incidences of psychedelic effects of ketamine like hallucination, dysphoria and psychotomimetic emergence reactions when ketamine is used in combination with a sedative-hypnotic (e.g., benzodiazepines, propofol) or a general anesthetic (e.g., halothane, nitrous oxide) is minimal or negligible. Hypnotic doses of propofol are reported to block these hallucinations. These findings are concordance with the study done by Arikani et al⁶ Brajesh et al⁷ Mahajan et al.¹⁵

Conclusion

It may be concluded from our present study that propofol with ketamine as an adjuvant in the dose of 0.75 mg/kg compared to propofol with fentanyl as adjuvant in the dose of 2 mcg/kg provides deep sedation. But propofol-ketamine group is more efficacious and provides better peri-operative

hemodynamic stability during anesthesia as compared to propofol-fentanyl group. Also the propofol-ketamine combination produces good analgesia with less requirement of rescue drug in post-operative period with fewer peri-operative complications than propofol-fentanyl combination. We have not encountered any psychotomimetic effects of injection Ketamine in the 0.75mg/kg dose. Hence propofol-ketamine combination is a better choice especially when hemodynamic stability is of great importance in patients undergoing dilatation and curettage.

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