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Effects on the Haemato-biochemical parameters by propofol, Ketofol and Etomidate as induction agent in Glycopyrrolate premedicated dogs maintained under Isoflurane anaesthesia

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Abstract

The present study was undertaken in 18 dogs with elective surgery (castration, spaying etc.) to evaluate the effect of propofol, Ketofol and etomidate as induction agents on Haemato-biochemical parameters in glycopyrrolate premedicated dogs maintained with isoflurane. The animals were randomly divided into three groups *viz*. group P, KP & E, comprising six animals in each group. All the animals of the three groups were premedicated with glycopyrrolate @ 0.01 mg/kg IM, 10 minutes prior induction. Propofol @ 6mg/kg IV, Ketofol @ 4mg/kg and etomidate @ 3mg/kg was administered as induction agent in group P, KP and E respectively. Maintenance of anaesthesia was carried out by using isoflurane in all the animals in all three groups. Parameters of Haemato-biochemical profile were studied at (baseline) 0 min and thereafter 5-, 15-, 30-, and 60-minutes time interval after induction.

Keywords: Glycopyrrolate, Propofol, Ketofol, etomidate, isoflurane, Heamato-biochemical parameters

Introduction

Propofol (2, 6-diisopropyl phenol) is an injectable anaesthetic agent belonging to the alkyl phenol group, with characteristics of very rapid redistribution from the brain to other tissues and can also efficiently eliminated from plasma through hydroxylation by one or more hepatic cytochrome P- 450 isoforms, results in a rapid onset of action, short duration of action with a complete and excitement-free rapid recovery, with good muscle relaxation, but with poor analgesic properties (Zoran et al., 1993, and Hall et al., 2001) ^[20, 5]. Ketofol administration offered effective sedation for spinal anaesthesia for gynaecologic, ophthalmologic and cardiovascular procedures in all age groups. The main advantage of this drug combination over alone propofol administration is the opposing hemodynamic and respiratory effects of each drug that enhance safety and efficacy and decrease the dose of propofol required for induction (Daabiss et al., 2009)^[3]. Association of ketamine and propofol may reduce unwanted adverse effects of both drugs, since these drugs act on different extremes - excitation and depression, respectively (Mair et al., 2009) ^[10]. Etomidate is a carboxylate imidazole derivative nonbarbiturate, short-acting, IV anaesthetic. Etomidate is characterized by better hemodynamic stability, minimal respiratory depression, and cerebral protective effects (Robert and Hiller, 2006) ^[15]. Etomidate induced minimal changes in cardiopulmonary function in hypovolaemic dogs (Pascoe et al., 1992)^[13]. This study was undertaken to evaluate effects of propofol, ketofol and etomidate on haemato-biochemical parameters as induction agent in glycopyrrolate premedicated dogs maintained under isoflurane anaesthesia.

Materials and Methods

The study was conducted on clinical cases of dogs of either sex those were brought to the Teaching veterinary Clinical Complex (TVCC), College of Veterinary Sciences and Animal Husbandry, Central Agricultural University, Aizawl, Mizoram. The animals were randomly divided into three groups *viz*. group P, KP & E, comprising six animals in each group. All the animals of the three groups were premedicated with glycopyrrolate @ 0.01 mg/kg IM, 10 minutes prior induction. Propofol @ 6mg/kg IV, Ketofol @ 4mg/kg and etomidate @ 3mg/kg was administered as induction agent in group P, KP and E respectively. Maintenance of anaesthesia was carried out by using isoflurane in all the animals in all three groups.

Haemato-biochemical parameters (Hb, PCV, TEC, TLC, Monocyte count, lymphocyte count, granulocyte count, serum glucose, ALT, AST, GGT, BUN, serum creatinine) were studied at (baseline) 0 min and thereafter 5, 15, 30, and 60 minutes after induction. Statistical analysis was carried out by SPSS version 20.

Results and discussion Haemato-biochemical parameters Haematological parameters

The mean ± SD values of haematological parameters, i.e., haemoglobin (g/dL), PCV (%), TEC (M/mm3), TLC (thousand/mm³), monocytes count (%), lymphocytes count (%), and granulocytes count (%) of different groups in different time intervals were observed and depicted in the Table 1. In this study non-significant decrease in Hb and PCV value was observed in group P, KP and E after administration of induction agents respectively and towards the end of the study gradual increase was observed in the Hb and PCV value towards the baseline in all three groups. Significant variations in Hb levels were observed between the groups but all the values within group and among groups did not exceed normal physiological limits. The mean values of PCV were recorded with non-significant variations at all (0, 5, 15, 30, 60 min) time intervals among all the groups. Similar findings for Hb and PCV were also observed by Bayan et al. (2002)^[2], Sharma et al. (2017) [16] and Shinde et al. (2018) [18] with propofol anaesthesia and significant decrease in Hb and PCV levels followed by gradual increase towards the end of study in propofol anaesthesia in dogs was observed by Thejasree et al. (2018)^[19]. Non-significant decrease in Hb and PCV were observed by Sharma et al. (2017)^[16] and Shinde et al. (2018) ^[18] and significant decrease followed by increase in Hb levels were observed by Thejasree et al. (2018) [19] in ketofol anaesthesia. Non-significant decrease in Hb but significant decrease in PCV value from baseline to end of trail was observed by Perk et al. (2002) [14] in etomidate/alfentanil anaesthesia in dogs. The decrease in Hb and PCV values might be due to the splenic pooling of circulating erythrocytes that occur with most of the anaesthetics or due to haemodilution in response to fluid therapy during anaesthetic protocol or due to inter compartmental fluid shift in order to maintain normal cardiac output Kumar et al. (2014)^[9] and Thejasree et al. (2018)^[19]. Non-significant decrease ($p \ge 0.05$) in TEC in group P initially followed by slight increase was observed towards the end of trial and similar findings with propofol anaesthesia were observed by Bayan et al. (2002)^[2], Sharma et al. (2017)^[16] and Shinde et al. (2018)^[17]. Highly significant ($p \le 0.01$) decrease in TEC values were recorded in group KP at different time intervals. The mean value of TEC showed significance difference $(p \le 0.01)$ among the groups but were within physiological range. However, in a study by Shinde 2018 [17] et al. non-significant decrease in TEC was observed in ketofol anaesthesia till the end of study in dogs where values were remained within normal physiological limits. In group E initially non-significant decrease in TEC followed by gradual increase towards the baseline at the end of study was observed but significant decrease in TEC value from baseline to the end of anaesthesia was observed by Perk et al. (2002)^[14] in etomidate/alfentanil anaesthesia in dogs. Significant difference of TEC was recorded between the three experimental groups at different time intervals till the end of the observation. Non-significant decrease $(p \ge 0.05)$ in TLC values in all three groups were observed initially followed by

slight increase in TLC at the end of anaesthetic trial. Similar pattern in TLC values was observed by Bayan et al. (2002)^[2] with propofol whereas Sharma et al. (2017)^[16] reported nonsignificant changes in TLC values with both propofol and Ketofol anaesthesia. Non-significant decrease in TLC value from baseline to the end of anaesthesia was observed by Perk et al. (2002)^[14] in etomidate/alfentanil anaesthesia in dogs. No significant difference of TLC was noted within the three experimental groups at different time intervals till the end of the observation. The decrease in both TEC and TLC values in this study might be due to individual variation in animals, or due to use of different anaesthesia agents in different groups or Splenic sequestration during anaesthesia might be the cause of decrease in TEC levels or combine effects of all. There was non-significant difference observed in monocyte count at different time intervals within all three groups but significant difference $(p \le 0.01)$ observed between group E compared group P and KP in the trail. No significant variations were recorded within the groups and between the groups. Sharma et al. (2017) ^[16] observed significant increase in monocyte in Ketofol group at the end of study compared to baseline whereas in propofol group non-significant decrease in monocyte count was observed at the end of study compared to baseline in group P non-significant decrease in monocyte was observed throughout the study period. Similar decreased in monocytes count in propofol anaesthesia was also observed by Nusory (2011)^[12]. In group E non-significant decrease in monocyte was observed from 15 min to the end of observation. The non- significant decrease in monocyte count might be due to splenic vasodilation due to anaesthetic drugs or might be due to administration of intravenous fluid administration throughout the anaesthetic protocol or might be combine both. Significant variation of monocyte count between groups might be due to higher baseline values of monocyte in group E or might be due to use of different anaesthetic drugs in different groups or both. There was no non-significant difference ($p \ge 0.05$) in lymphocyte count was observed throughout the anaesthetic protocol among the groups and between the groups and all values are within normal physiological range although gradual decrease in lymphocyte value was observed in all three groups. Similar findings of decreased in lymphocytes count in propofol anaesthesia was observed by Kelawala et al. (1996)^[8] and Surbhi (2008)^[18] whereas Sharma et al. (2017)^[16] observed non-significant decrease in lymphocyte in both propofol and ketofol group initially but at the later stage non-significant increase in lymphocyte count. In group E, at 5 min nonsignificant increase in lymphocyte count followed by gradual decrease till the end of study period was observed. The nonsignificant decrease in lymphocyte count might be due to splenic vasodilation due to anaesthetic drugs or might be due to administration of intravenous fluid administration throughout the anaesthetic protocol or might be combine both. Non-significant increase $(p \ge 0.05)$ in granulocyte count was observed throughout the anaesthetic protocol in both group P and KP whereas in group E initially non-significant decrease $(p \ge 0.05)$ in granulocyte count observed followed by gradual increase in lymphocyte count was observed till the end of anaesthesia. All the variations remain within normal physiological range. The mean values of granulocyte count were recorded with slight non-significant variations throughout the trail among all three groups. Similar findings of increased in neutrophil count in propofol anaesthesia was recorded by Kelawala et al. (1996)^[8] and Surbhi (2008)^[18]

whereas, Sharma et al. (2017) [16] observed non-significant increase in neutrophil in propofol group initially but gradual decrease towards the baseline and goes down below the baseline level. In a relatively similar study by Sharma et al. (2017) ^[16] non-significant increase in neutrophil count was observed in Ketofol anaesthesia and remained higher than the baseline value at the end of anaesthesia. In group E initially non-significant decrease in granulocyte at the initial stage followed by gradual increase in granulocyte was observed. No significant variations were recorded within the groups and between the groups. Increase in overall granulocytes might be due to increase in neutrophil count. The non- significant increase in granulocytes count might be due to splenic vasodilation due to anaesthetic drugs or might be due to administration of intravenous fluid administration throughout the anaesthetic protocol or Surgical and anaesthetic stress leading to stimulation of the adrenal cortex or might be combination of all factors.

2. Biochemical parameter

The mean \pm SD values of bio chemical parameters, i.e., serum glucose (mg/dL), ALT (U/L), AST (U/L), GGT (U/L), BUN (mg/dL), and serum creatinine (mg/dl) of different groups in different time intervals were observed and depicted in the Table 2. In all the group glucose value significantly ($p \le 0.05$) & $(p \le 0.01)$ increased till the end of observation but changes remain within normal physiological range. Among the groups significantly increase glucose value was recorded in group P and group KP at 30- and 60-min time interval as compared to group E. Similar increase in blood glucose was observed by Bayan (2002)^[2] with propofol and Anandmay et al. (2016)^[1] with buprenorphine- propofol and alone propofol anaesthesia at 1 hrs. post induction in canine. In group KP significant increase in glucose value was observed in 30- and 60-min time interval and similar significant increase in blood glucose in Ketofol induced dogs was recorded by Njoku (2015)^[11]. However non-significant increase in blood glucose value with propofol at 60 min and Ketofol at 60 min was observed by Sharma et al. (2018). In group E significance difference was observed at the end period of the study and Similar significant findings was observed by Perk et al. (2002) [14] with etomidate/alfentanil anaesthesia in dogs. Sumer et al. (2012) and Kaushal et al. (2015) [7] observed increase in glucose values in the patients treated with etomidate. The increase was observed in the present study might be due to cortisol and catecholamine mediated gluconeogenesis along with decrease peripheral use of glucose reported by Bayan et al. (2002)^[2]. There was no significant difference ($p \ge 0.05$) in ALT and AST values were observed throughout the anaesthetic trials among the groups and between the groups but non-significant increase in ALT values was observed in all three groups throughout the study period but the values remained within normal physiological range. In group P non-significant increase in ALT and AST was observed throughout the study period and similar findings of non-significant increase in ALT and AST were observed by Thejasree et al. (2018) [19] and Shinde et al. (2018)^[17] in propofol anaesthesia. In group KP non-significant increase in ALT and AST were observed throughout the study period, similar to the findings of Thejasree et al. (2018)^[19]. However, non-significant decrease in ALT and AST were observe in ketofol group in dogs by Shinde et al. (2018)^[17] which was contrary to the present findings. Non-significant increase in ALT and AST were observed up to 60 min compared to the baseline and non-

significant increase in ALT and AST was observed by Hareesh et al. (2018)^[6] in etomidate anesthesia, which was similar to the present finding. Perk et al. (2002)^[14] observed non-significant decrease in ALT and non-significant increase in AST in etomidate/alfentanil anaesthesia in dogs. Highly significant difference ($p \le 0.01$) in GGT values were recorded in group P and group KP at different time interval except 0 min in compared to group E. GGT values were significantly increased up to 30 min. compared to baseline followed by decreased end of the experiment. These findings are similar to the findings of Ward et al. (2006) in captive wild dogs and Nusory (2011)^[12] during propofol anaesthesia in dogs and Anandmay et al. (2016)^[1] observed non-significant increase in GGT values at 1 hour interval was recorded compared to baseline in propofol anaesthesia in dogs. Significant difference in GGT values within the groups were recorded throughout the experiment except at the baseline. GGT is more sensitive than ALT and AST for detecting liver damage. The transient variations in ALT, AST and GGT values in the present study were within normal physiological limits might be indicative of non-toxic/less harmful effect of all the anaesthetic drugs on hepatobiliary system. In group P, KP and E non-significant ($p \ge 0.05$) increase in BUN values were observed till 30 min. but towards the end part of observation it declined towards the baseline. There were highly significant $(p \le 0.01)$ variations in the mean values of BUN were recorded among the groups at different time interval but remain normal physiological range. Non-significant increase in BUN values were observed in group P and KP up to 30 min followed by decrease in BUN value towards the baseline towards the end of the study. Similar findings were observed by Sharma et al. (2017) ^[16] and Shinde et al. (2018) ^[17] with propofol. However, Shinde et al. (2018)^[17] observed decrease in BUN values after recovery compared to baseline in dogs in ketofol anaesthesia. In group E, non-significant increase followed by decrease in BUN values were observed in the study period but all were remained within normal physiological limit. Perk et al. (2002) ^[14] observed decrease in BUN values with etomidate/alfentanil anaesthesia in dogs. BUN values recorded at baseline and all other time intervals showed significant variations between all three groups but were within the normal range. In both group P and KP highly significant $(p \le 0.01)$ increase in creatinine value was observed throughout the study period compared to baseline (0 min) but values remained within normal physiological range. In this study significant increase in the creatinine values were observed in group P up to 30 min and decreases towards the baseline at the end part of study and relatively similar findings were observed by Sharma et al. (2017)^[16] and Shinde et al. (2018) ^[17] observed non-significant increase in creatinine during maintenance followed by decrease after recovery from anaesthesia. In group KP increase in the creatinine values were observed up to at the end of anaesthesia and similar findings were observed by Sharma et al. (2017) [16] where increase in creatinine was observed at the end of anaesthesia in Ketofol group compared to baseline. In group-E, creatinine values were non significantly increased than its base value during anaesthetic trial at different time intervals. It fluctuated within normal physiological limits. Its might be due to etomidate does not affect the renal blood flow and glomerular filtration rate recorded by Grimm et al. (2015)^[4]. The variations in BUN and creatinine in all groups in different time intervals might be due to use of different induction agents in different groups and random selection of animals in

groups or might be both. Increase in BUN and creatinine value during trail period in all groups might be due to the

temporary inhibitory effects of anaesthetic drugs on the renal blood flow leading to decrease in glomerular filtration.

 Table 1: Mean ± SD values of different haematological parameters at different time intervals in group P, KP and E Values in the same row with different superscripts (small front) differ significantly and Values in the same column with different superscripts (capital front) differ significantly and Values in the same column with different superscripts (capital front) differ significantly

Parameters	Groups	Baseline (0 min)	5 min	15 min	30 min	60 min	Significance
НВ	Р	14.17 ± 0.99^{Ba}	13.11 ± 1.05^{Ba}	12.72 ± 1.08^{Ba}	12.97 ± 1.03^{Ba}	13.37 ± 0.87^{Ba}	NS
	KP	$14.91 \pm 1.4^{\text{Ba}}$	14.29 ± 1.46^{Ba}	13.68 ± 1.49^{Ba}	13.30 ± 1.44^{Ba}	13.73 ± 1.48^{Ba}	NS
	Е	12.24 ± 1.03^{Aa}	11.56 ± 1.21^{Aa}	11.16 ± 1.08^{Aa}	11.46 ± 0.91^{Aa}	11.78 ± 0.95^{Aa}	NS
	Significance	**	**	**	*	*	
PCV	Р	$49.18\pm5.49Aa$	46.92 ± 5.50 Aa	46.60 ± 5.61 Aa	47.19 ± 5.71Aa	47.94 ± 5.57Aa	NS
	KP	$45.97 \pm 3.24 Aa$	44.48 ± 3.53 Aa	43.95 ± 3.26 Aa	43.40 ± 3.51 Aa	44.00 ± 3.41 Aa	NS
	E	$45.21 \pm 2.39 Aa$	42.96 ± 1.77 Aa	42.88 ± 2.42 Aa	43.91 ± 2.72 Aa	44.68 ± 2.80 Aa	NS
	Significance	NS	NS	NS	NS	NS	
TEC	Р	7.19 ± 0.48^{Ba}	6.67 ± 0.60^{Ba}	6.52 ± 0.64^{Ba}	6.68 ± 0.69^{Ba}	6.90 ± 0.68^{Ba}	NS
	KP	7.23 ± 0.39^{Bb}	6.76 ± 0.34^{Ba}	6.33 ± 0.56^{Ba}	6.25 ± 0.48^{Ba}	6.53 ± 0.48^{Ba}	*
	E	6.01 ± 0.84^{Aa}	5.29 ± 0.61^{Aa}	5.27 ± 0.45^{Aa}	5.39 ± 0.50 Aa	5.65 ± 0.49^{Aa}	NS
	Significance	**	**	**	**	**	
TLC	Р	$12.65 \pm 1.36 \text{Aa}$	$11.19 \pm 1.05 Aa$	10.66 ± 1.19 Aa	$11.07 \pm 1.47 Aa$	11.68 ± 1.43 Aa	NS
	KP	$12.63 \pm 1.70 Aa$	$12.06 \pm 1.71 \text{Aa}$	11.68 ± 1.78 Aa	11.33 ± 1.72 Aa	11.72 ± 1.73 Aa	NS
	E	15.05 ± 2.2 Aa	13.47 ± 2.39 Aa	13.45 ± 2.06 Aa	$13.60 \pm 2.50 \text{Aa}$	13.90 ± 2.38 Aa	NS
	Significance	NS	NS	NS	NS	NS	
Monocyte count	Р	4.28 ± 0.77^{Aa}	4.08 ± 0.75^{ABa}	3.91 ± 0.82^{ABa}	3.73 ± 0.80^{Aa}	3.66 ± 0.78^{Aa}	NS
	KP	3.78 ± 0.60^{Aa}	3.50 ± 0.49^{Aa}	3.25 ± 0.40^{Aa}	3.11 ± 0.37^{Aa}	3.05 ± 0.39^{Aa}	NS
	E	4.78 ± 0.74^{Aa}	4.78 ± 0.61^{Ba}	4.65 ± 0.53^{Ba}	4.60 ± 0.39^{Ba}	4.55 ± 0.52^{Ba}	NS
	Significance	NS	**	**	**	**	
Lymphocyte count	Р	$18.11\pm2.67Aa$	17.51 ± 2.45 Aa	17.13 ± 2.54 Aa	$16.91 \pm 2.34 Aa$	$16.58 \pm 2.38 Aa$	NS
	KP	$17.70 \pm 1.94 Aa$	$17.04 \pm 1.84 Aa$	$16.52 \pm 1.71 \text{Aa}$	$16.10\pm1.67Aa$	$15.66 \pm 1.64 Aa$	NS
	E	$18.53 \pm 2.35 Aa$	18.81 ±2.62Aa	$18.63 \pm 2.38 \text{Aa}$	$17.98 \pm 1.95 \text{Aa}$	$17.68 \pm 1.89 Aa$	NS
	Significance	NS	NS	NS	NS	NS	
Granulocyte count	Р	$77.60 \pm 2.75 Aa$	$78.43 \pm 2.59 \text{Aa}$	$78.95 \pm 2.74 \mathrm{Aa}$	79.40 ± 2.53 Aa	79.75 ± 2.67 Aa	NS
	KP	$78.\overline{51} \pm 2.16 \text{Aa}$	79.45 ± 1.97 Aa	80.22 ± 1.78 Aa	$80.\overline{78} \pm 1.\overline{74}Aa$	$81.\overline{28 \pm 1.63}$ Aa	NS
	E	$76.\overline{68 \pm 2.77}$ Aa	76.40 ± 2.76 Aa	76.71±2.41Aa	77.41 ± 2.18 Aa	79.56 ± 2.22 Aa	NS
	Significance	NS	NS	NS	NS	NS	

 Table 2: Mean ± SD values of different biochemical parameters at different time intervals in group P, KP and E Values in the same row with different superscripts (small front) differ significantly and Values in the same column with different superscripts (capital front) differ significantly

Parameters	Groups	Baseline (0 min)	5 min	15 min	30 min	60 min	Significance
Serum Glucose	Р	89.51 ± 11.23^{Aa}	95.37 ± 11.27^{Aa}	98.90 ± 10.82^{Aab}	$103.32 \pm 10.50^{\text{Bab}}$	$110.15 \pm 9.89^{\text{Bb}}$	*
	KP	92.71 ± 9.28^{Aa}	97.38 ± 8.44^{Aab}	102.18 ± 7.46^{Aab}	107.16 ± 7.38^{Bb}	117.86 ± 7.47^{Bc}	**
	Е	84.7 ± 6.29^{Aa}	87.50 ± 5.68^{Aab}	90.03 ± 5.60^{Aab}	92.58 ± 4.90^{Aab}	98.30 ± 4.34^{Ab}	*
	Significance	NS	NS	NS	*	**	
ALT	Р	$29.80 \pm 2.63 Aa$	30.41 ± 2.76 Aa	$31.38 \pm 2.92 Aa$	31.93 ±.33Aa	$32.44 \pm 3.16 \mathrm{Aa}$	NS
	KP	$29.84 \pm 1.05 Aa$	30.43 ± 1.12 Aa	31.15 ± 1.2 Aa	$3180 \pm 1.15 Aa$	32.20 ± 1.15 Aa	NS
	E	$29.71 \pm 2.42 Aa$	29.98 ± 2.49 Aa	$30.50\pm2.43Aa$	30.71 ± 2.66 Aa	$31.2\pm3.00Aa$	NS
	Significance	NS	NS	NS	NS	NS	
AST	Р	16.72 ± 1.66 Aa	$16.97 \pm 1.66 Aa$	$17.25 \pm 1.60 Aa$	17.29±1.67Aa	17.6 ± 1.70 Aa	NS
	KP	$16.50\pm1.62Aa$	$16.87 \pm 1.57 Aa$	$17.23 \pm 1.52 Aa$	$17.35 \pm 1.57 Aa$	17.68 ± 1.55 Aa	NS
	E	$16.56 \pm 1.61 Aa$	16.87 ± 1.42 Aa	$17.20 \pm 1.32 Aa$	$17.36 \pm 1.58 Aa$	17.60 ± 1.56 Aa	NS
	Significance	NS	NS	NS	NS	NS	
GGT	Р	$2.87\pm0.41 Aa$	$3.07\pm0.37Ba$	$3.19\pm0.34Bab$	$3.62\pm0.39Bb$	$3.29 \pm 0.36 Bab$	*
	KP	2.90 ± 0.23 Aa	$3.14\pm0.18Ba$	$3.44\pm0.23Bb$	$3.73 \pm 0.20 Bc$	3.53 ± 0.27 Bbc	**
	Е	$2.50b\pm0.30Aa$	$2.53\pm0.22Aa$	$2.57\pm0.26\mathrm{Aa}$	$2.60\pm0.23Aa$	$2.75\pm0.27Aa$	NS
	Significance	NS	**	**	**	**	
BUN	Р	$10.01\pm0.96Ba$	$10.38 \pm 1.10 Ba$	$10.65\pm1.02Ba$	$11.25\pm1.02Ba$	$10.82 \pm 1.06 Ba$	NS
	KP	$11.36\pm0.48Ca$	$11.68\pm0.49Ca$	$12.13\pm0.47Ca$	$12.06\pm0.59Ba$	$11.98 \pm 0.44 Ca$	NS
	E	$8.00\pm0.94Aa$	$8.03 \pm 1.00 Aa$	$8.05\pm0.87Aa$	$8.20\pm0.93Aa$	8.14 ± 0.89 Aa	NS
	Significance	**	**	**	**	**	
Serum Creatinine	Р	$0.94\pm0.07Ba$	$1.04\pm0.08Bab$	$1.16\pm0.12Bbc$	$1.26\pm0.08Bc$	$1.10\pm0.14Ab$	**
	KP	1.03 ± 0.11 Ba	$1.11\pm0.10Ba$	$1.25\pm0.08Bb$	$1.30\pm0.09Bbc$	$1.40\pm0.14Bc$	**
	Е	0.79 ± 0.13 Aa	$0.84\pm0.12Aa$	0.87 ± 0.15 Aa	0.90 ± 0.16 Aa	$0.9\overline{4\pm0.13}Aa$	NS
	Significance	**	**	**	**	**	

Conclusion

The haemoglobin (Hb), packed cell volume (PCV), total leukocyte count (TLC), differential leukocyte count (DLC), lymphocyte count and granulocyte count did not show any significant difference in all three groups. But total erythrocyte count (TEC) showed significant decrease in group KP but non-significant variations of TEC were observed in group P and E. The blood glucose showed significant increase in all three groups. ALT, AST, and BUN did not show any significant difference in three groups. In group P and KP significant increase in GGT and creatinine was observed where as non-significant changes in GGT and creatinine was observed in group E. Based on the analysis of Haematobiochemical parameters it was found that etomidate have slight advantage over propofol and Ketofol as a better induction agent but overall evaluation based on clinical, anaesthetic, cardiopulmonary, and Haemato-biochemical parameters, it was found that all three induction agents are relatively safe in glycopyrrolate premedicated dogs maintained under isoflurane anaesthesia.

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