# Comparison of midazolam and propofol during induction in elective CABG

Bilge ÇELEBİOGLU', Ömür ERÇELEN', İlhan PAŞAOĞLU', Alev GÖL', Ülkü AYPAR', Kemal ERDEM'

Depts. of Anaesthesiology', and Cardivascular Surgery', Hacettepe University, Ankara, TURKEY

Hemodynamic changes during induction of anaesthesia, with midazolam and propofol were compared in patients undergoing elective CABG.

0.2 mg/kg/IV and Group Group M (Midazolam) was administered midazolam P(propofol) was mg/kg/IV. All patients were adndnistered fentanyl 2 mg/kg/hr/IV. together with 50 % N20, 50 % 02 after induction. systolic arterial pressure (SAP), diastolic arterial pressure measured heart rate (HR). (DAP). (MAP), central venous pressure (CVP), before induction. after midazolam or propofol. after fentanyl, postintubation and after sternotomy. In both groups, the total dose of fentanyl was similar.

There was no difference in these parameters between midazolam and propofol. As a result this study shows that for the induction of anesthesia midazolam has no advantage over propofol considering the cardiac parameters. [Turk J Med Res 1994; 12(2): 66-69]

KeyWords: Anesthesia, Midazolam, Propofol.

Midazolam is a water-soluble benzodiazepine compound that can be used for sedation and as a preanaesthetic medication and an anaesthetic adjuvant (1,2). Midazolam causes only minor haemodynamic changes in surgical patients without concomitant debilitating affections. It has been considered as being safe in cardiac disease: its nitroglycerin-like vasodilating effect is benfical both in case of pulmonary hypertansion and congestive heart failure, while cardiac output is unaffected in patients with ischaemic heart disease (3,4).

Propofol is the most recently introduced parenteral anaesthetic agent and because of this data concerning its availability for cardiac surgery are limited. The adverse hemodynamic effects of propofol when given bolus dose correlate with overshoot in blood concentration. If used appropriately propofol may have significant advantages in cardiac anaesthesia (5).

In this study, we compared the changes of cardiac parameters during induction with the midazolam and propofol.

Correspondence: Bilge CELEBIOGLU
Hale Sok. 12/4 Maltepe
06570, Ankara, TURKFY

#### PATIENTS AND METHODS

The study included 24 patients of ASA grade II-III (over the EF 60 %), scheduled for CABG surgery. Patients were selected randomly to two groups: midazolam (group M) IV and propofol (group P) IV.

Group M: Anaesthesia was induced by administration of midazolam 0.2 mg/kg/IV and after 5 minutes subsequently maintained with 50 % 02 and 50% N20 and fentanyl 2 ug/kg/hr/IV.

Group P: Anaesthesia was induced by administration of propofol 2 mg/kg/IV and after 5 minutes anaesthesia was maintained like group M.

In both groups one minute after the initial dose of fentanyl, vecuronium 0.1 mg/kg/IV was given. After the development of complete muscle relaxation tracheal intubation was made when required intermittent doses of vecuronium were given throughout anaesthesia based on the response of the thumb to ulnar nerve stimulation (ProMed-Easistim, Lot. No. 0.02).

Haemodynamic data were collected and evaluated at the following times before and throughout anaesthesia.

- ${\hbox{1. Preinduction: before IV midazolam and propofol}} \\ {\hbox{administration.}}$
- 2. Postmidazolam and postpropofol but prefentanyl: before fentanly IV.

Table 1. The demographic data of patients

	Midazolam	Propofol
Weight (kg) Height (cm) Age (year)	63.3±3.7 164.83+2.3 60.42+2.5	72±2.2 168.17+1.7 58.33±1.8
Male	10±0	12±0
Sex Female	2+0	0±0
Ejection Fraction E F %	60.92±2.4	62.5±2.9

Table 2. The cardiac data of patients

		Midazolam	Propofol
Pre	HRb/m	67.7+3.7	64.17±2.7
Induction	SAPmmHg	147.6±7.5	164.8±7.6
	DAP mmHg	73.2±3.7	77.7±3.4
	MAPmmHg	103.3±49	107.7±3.8
After	HRb/m	69.6±3.9	72.5±4.2
Midazolam	SAPmmHg	124.0±6.7	132.8±10
Propofol	DAP mmHg	62.8±3.4	66.2+4.4
	MAPmmHg	86.1 ±4.5	91.7±5.8
After	HRb/m	63.0±3.3	70.58±2.6
Fentanyl	SAPmmHg	107.1 ±1.7	111.2±9.7
	DAP mmHg	58.8+3.7	59.7±5.1
	MAPmmHg	76.7+4.3	75.3+6.2
Post	HRb/m	64.6±4.5	67.0+3.1
Intubation	SAPmmHg	129.1+9.7	123.7+6.3
	DAP mmHg	71.3±5.4	68.4±3.4
	MAP mmHg	80.7±3.1	86.9±4.2
Post	HRb/m	66.1 ±3.2	63.0±3.2
Sternotomy	SAPmmHg	121 <b>.318</b> .8	115.7+4.3
	DAP mmHg	63.9+4.1	65.25±2.9
	MAPmmHg	86.4±5.9	83.3+3.4
Peroperatif			
	CVP mm Hg	7.1 ±2.1	7.2±1.8

- 3. Postfentanyl but preintuba'.ion: 5. minutes after fentanyl and vecuronium IV injection.
  - 4. After sternotomy.
- \ Cardiac parameters evaluated were
  - Heart rate (HR) beats/minute
- Systolic systemic blood pressure (SAP)  $_{\rm mmHg}$
- Diastolic systemic blood pressure (DAP)  $_{\rm mmHg}$ 
  - Mean systemic blood pressure (MAP) mmHg
  - Central venuos pressure (CVP) mmHg

The cardiac parameters were evaluated for the two groups at each sampling time during the procedure using the Student's test. Statistical significance was defined at p<0.05. There were no significant dif-

ference in all parameters between midazolam and propofol.

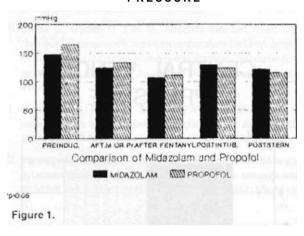
#### RESULTS

Intravenous injection of midazolam and propofol caused no significant changes on haemodynamic variables

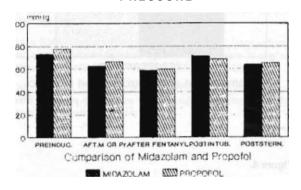
SAP changes were shown in Figure 1, DAP changes were shown in Figure 2, MAP changes were shown in Figure 3, CVP changes were shown in Figure 4, HR changes were shown in Figure 5.

They were showed no stastically significant changes between two groups.

## SYSTOLIC ARTERIAL PRESSURE



## DIASTOLIC ARTERIAL



10:0.05

Figure 2.

#### DISCUSSION

Several intravenous induction agents including thiopentone, fentanyl, alfentanil, midazolam and propofol have

Turk J Med Res 1994; 12 (2)

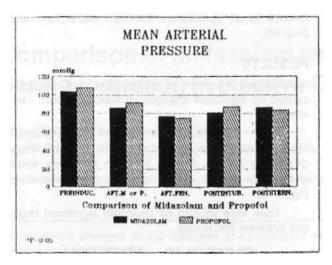
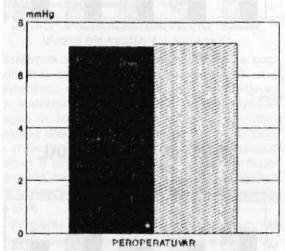


Figure 3.

## CENTRAL VENOUS PRESSURE



Comparison of Midazolam and Propofol

p-a.oa

Figure 4.

been given at induction of anaesthesia (6-10). There has been a conflict of views regarding the suitability of midazolam as an intravenous induction agent. Some authors believe that midazolam is an unreliable induction agent while others find it acceptable for induction (11).

Propofol has been very popular as an induction agent since its introduction as an emulsion preparation

(5). Propofol is the most recently introduced intravenous anaesthetic agent and data about its suitability for cardiac surgery are limited. Initial studie examined propofol as an induction agent in the presence of coronary artery disease noted a mean reduction of 28 % in systolic blood pressure related primarily to a 25 % fall in CVP (12). An alternative approach has been to avoid the use of propofol for induction of anaesthesia (13). Vermeyen et al. also noted good protection from the haemodynamic response to sternotomy using a combination of moderate dose fentam and propofol (12).

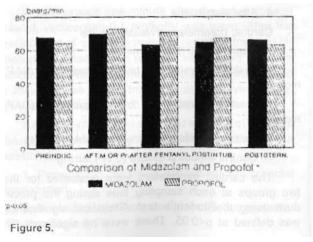
When midazolam is administered in combination with fentanyl, hypotension secondary to myocardial depression may be seen (14). This effect can be limited by slow administration end careful titration of the drug to the required and point and this has become a popular induction technique in cardiac anaesthesia (15). Further studies are required to assess the potential benefits of midazolam and the newer opioids for cardiac anaesthesia (15).

In some studies, the free plasma concentration of midazolam and propofol were compared and no statistically significant alteration were found (16).

In our study, we administered midazolam-fentanyl and propofol-fentanyl combination for induction in coronary artery surgery. Between two groups no significant changes has been found about all parameters. The safety of midazolam-fentanyl and propofol-fentanyl in anaesthetic induction for induction followed by maintenance of analgesia with fentanyl-50% N20 - 50 % 02 is also confirmed by the lack of adverse occurences intraoperatively in these patients such as hypo or hypertension, brady-or tachycardia, arrhytmias etc. Therefore, no vasoactive drugs, inotropic agents or p-blockers were needed throughout the induction of anaesthesia for the routine nitroglycerin IV infusion.

In conclusion, midazolam-fentanyl or propofol-fentanyl induction/intubation sequence caused no adverse

#### HEART RATE



< Turk J Med Res 1994; 12 (2)

haemodynamic changes in coronary artery surgery in the doses and sequence unitilized in our study. We thought that midazolam-fentanyl and propofol-tentanyl combination can be used for induction of anaesthesia in coronary artery surgery. But, midazolam doesn't have any advantage when compared with propofol.

#### Elektif koroner arter operasyonlarında indüksiyon sırasında midazolam ve propofol kullanımının karşılaştırılması

Çalışmamızda, elektif CABG hastalarda. geçiren midazolam propofol ile yapılan anestezi indüksivonu sırasında oluşan hemodinamik değişiklikler karşıla ştırıldı.

Grup М (midazolam)'daki hastalara indüksivon SIrasında0 2 mgjkg/iv midazolam. Gnup P(propofol)'daki hastalara ise 2 mg/kg/ivpropofol uygulan-Bütün hastalara anestezi idamesinde 2 mg/kg/saat/iv hızda fentanil infüzvonu ve %50 N20. %50 02 kullanıldı. Kalp atım hızı. sistolik. diastolik. ortalama arter basıncları. santral venöz basınç deăisiklikleri:

1indüksivon önücesi2- midazolam veya propofol 3-fentanil veriliminden sonra sonrası 4-intübasyon 5-sternotomi sonrası takip edildi. sonrası Her iki grupta ölçülen parametrelerden elde edilen soistatistiksel olarak anlamlı bir fark nuçlar arasında saptanamadı (p>0.05).Ви çalışmanın sonucunda, CABG sırasında. anestezi indüksiyonunda, midazolamın propofole oranla bir üstünlüğü olmadığı kavarılabilinir. [Turk J Med Res 1994; 12(2): 66-69]

#### REFERENCES

- Richard I Hall, Ian M. Schwieger, Carl C, et al. The anesthetic efficacy of midazolam in the enfluorane-anesthetized dog. Anesthesiology 1988; 68: 862-6.
- Fragen RJ, Gahl F, Caldwell N. A water-soluble benzodiazepine. Ro 21-3981, for induction of anaesthesia. Anasthesiology 1978; 49:41-3.
- Healy TEJ, Kay B. Complement: Total intravenous anaesthesia. Elseiver. Amsterdam, New York. IOxford 1991; 21: 158-9.

- Fragen RJ, Meyers SN, Barresi V, et al. Hemodynamic effects of midazolam in cardiac patients. Anesthesiology 1979: 51:103
- Sebel PS, Lowdon JD. Propofol: a new intravenous anesthetic. Anesthesiology 1989; 71: 260-77.
- Short TG, Gallethy DC, Plummer JL The hypnotic and anaesthetic action of thiopenthone and midazolam alone and in combination. British Journal of Anaesthesia. 1991; 66: 13-9.
- Tverskoy M, Fleyshman G, Bradley EL, et al. Midazolamthiopental anesthetic interaction in patients. Anesthesia and Analgesia. 1988; 67: 342-5.
- Ben-Shlomo I, Abd-el-Khalim H, Ezry J, et al. Midazolam acts synergistically with fentanyl for induction of anaesthesia. Br. Journal of Anaesthesia 1990; 64: 45-7.
- Vinik HR, Bradley EL, Kissin I. Midazolam-alfentanil synergism for anesthetic induction in patients. Anesthesia and Analgesia 1989; 69: 213-7.
- Cross G, Gylard D, Lim M. Atropine induced heart rate changes: a comparison between midazolam-fentanyl-propofol-N20 and midazolam-fentanyl-thiopentone-enflurane-N20 anaesthesia. Canadian Journal of Anaesthesia 1990; 37: 416.0
- Gamble JAS, Kawar P, Dundee JW, et al. Evaluation of midazolam as an intravenous induction agent. Anaesthesia 1981; 36:868-73.
- Vermeyen KM, Erpels FA, Janssen LA, et al. Propofol-fentanyl anaesthesia for coronary bypass surgery in patients with good left ventricular function. Br J Anaesth. 1987; 59: 1115-20.
- Russel GN, Wright EL, Fox MA, et al. Propofol-fentanyl anaesthesia for coronary artery surgery and cardiopulmonary bypass. Anaesthesia 1989; 44: 205-8.
- Heikkila H, Jalonen J, Arola M, et al. Midazolam as an adjunct to high dose fentanyl anaesthesia for coronary artery bypass grafting operations. Acta Anaesthesiol. Scand. 1984; 28: 683-9.
- Raza SMA, Masters RW, Vasireddy AR et al. Haemodynamic stability with midazolam-sulfentanil analgesia in cardiac surgical patients. Can J. Anaesth. 988; 35: 518-25.
- Teh J, Short TG, Wong J, et al. Pharmocokinetic interactions between midazolam and propofol: an infusion study. Br. J. of Anaesthesia 1994: 2: 62-5.