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INTRODUCTION: Desflurane is an investigational, yolatile anesthetic with a blood-gas partition coefficient of 0.421 and a MAC of approximately 6.0 %. This open-label, randomized trial compared desflurane (DES) and isoflurane (ISO) during

elective coronary artery surgery (CAS).

METHODS: With institutional approval and written informed consent, we studied 17 patients with ejection fractions (EF) ≥ 0.35. Premedication consisted of the patient's usual antianginal drugs, plus morphine 0.1 mg/kg IM, and scopolamine 6µg/kg IM. After insertion of monitoring catheters, anesthesia was induced with IV thiopental 2 mg/kg/min until loss of the lash reflex. Pancuronium 0.1 mg/kg and fentanyl 10μ g/kg were then infused over 5 min, and \leq 1 MAC end-tidal (ET) DES or ISO administered in 100% O₂. After intubation, ventilation was controlled to achieve an ETCO₂ of 35 mmHg. The inhaled agent, ≤ 2 MAC, was titrated to maintain heart rate (HR) ≤ 120% and mean systemic arterial pressure (MAP) ≥ 70% and ≤ 120% of ward values. If the inhaled agent was ineffective, or if myocardial dysfunction or ischemia occurred, additional fentanyl, and/or vasoactive drugs were given. Systemic arterial, pulmonary arterial, and central venous pressures, ECG leads II and CS5, ST segment trends and ET agent concentration (AC) were continuously monitored and recorded. Thermodilution cardiac output (CO), and wedge pressure (PCWP), were recorded at baseline (B), and 1 min after induction (IND),

A322

TITLE: EFFECT OF N2O ON VITAMIN B12 AND FOLATE METABOLISM IN PATIENTS

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Patients occasionally develop hematologi-Introduction: cal and neurological abnormalities after surgery with an anesthetic technique that includes N2O. These abnormalities presumably result from N2O-inactivation of the vitamin B12-dependent enzyme methionine synthase and disruption of vitamin B12-folate interrelationships. We designed this study to measure the effect of N2O on vitamin B₁₂ and folate status in two groups of patients: 1) elderly patients having total hip replacements (HIP), and 2) patients undergoing resection of acoustic neuromas (AN) requiring prolonged anesthesia and surgery. Methods: With approval from our committee on human research and informed consent, we studied 49 patients (23 HIP, 26 AN). All patients received isoflurane, but were randomly assigned to receive either N2O 50-70% or O2 Inhaled isoflurane concentrations were adjusted to maintain clinically acceptable levels of anesthesia. Serum vitamin B₁₂ and serum and red cell folate were determined by radioassay preoperatively and on postoperative day 1. Spot urine samples were collected preoperatively and on postoperative days 1-3. Urine was assayed by spectrophotometric techniques for formic acid and formiminoglutamic acid (FIGLU); the excretion of

intubation (INT), and sternotomy (S). Serial 12-lead ECG and CK-MB concentrations were measured postoperatively. Statistical analysis employed t-test, Fisher Exact Test or ANOVA when appropriate.

RESULTS: Eight patients received DES and 9 ISO. The groups did not differ with respect to demographics, preoperative hemodynamics or EF. Prebypass hemodynamics were similar in both groups (Table, Mean \pm SD), as were the respective (DES vs ISO) mean ET-AC/MAC (0.68 \pm 0.19 vs 0.68 \pm 0.12), the number of patients given additional fentanyl (2 vs 2), propranolol (2 vs 2) and vasodilators (1 vs 1). New prebypass ST depression \geq 0.1 mV (2 vs 2), initial postoperative CK-MB (48±34 vs 56±34 IU) and the incidence of new Q waves (0 vs 0) were not different.

CONCLUSION: Desflurane is comparable to isoflurane when used as the primary anesthetic agent for CAS.

Variable	Agent	Base	Ind	Int	Stern
HR	Des	54±13	57±8	61±6	63±11
(b/min)	Iso	59±14	59±13	71±18	68±10
MAP	Des	98±16	73±9	76±9	89±10
(mmHg)	Iso	94±9	72±15	80±11	99±10
CI	Des	2.4±.6	2.2±.4	2.3±.3	2.1±.4
(l/min/m ²)	Iso	2.9±.8	2.3±.6	2.7±.6	2.4±.4
PCWP	Des	17±5	12±2	11±2	17±3
(mmHg)	Iso	15±4	12±3	11±3	14±5

REFERENCE: 1. Anesth Analg 66: 971-973, 1987.

these compounds in urine may be markedly elevated in individuals with a folate deficiency. Data were examined as mean values and change and percentage change from preoperative values using paired or unpaired t-tests, the Mann-Whitney test, or Fisher's exact test. Results: Average duration of anesthesia was 3.3 hrs in

HIP and 9.6 hrs in AN patients. No patient was deficient in vitamin B₁₂ and vitamin B₁₂ levels did not change on postoperative day 1 in either those exposed or not exposed to N2O. Prior to surgery, no patient had an abnormally low serum folate, although 19 patients had low red cell folate values. Serum folate levels tended to increase on the first postoperative day in patients exposed to N2O, but decreased in those not exposed. Red cell folate levels increased on postoperative day 1, independent of exposure to N2O. For HIP patients, ratios of formic acid-tocreatinine and FIGLU-to-creatinine in urine did not differ significantly between patients anesthetized with and without N2O. For AN patients given N2O, ratios of FIGLUto-creatinine in urine increased during and immediately after anesthesia when compared to preoperative values (p=0.0057), whereas patients not given N2O had FIGLU-tocreatinine ratios that were lower than preoperative values. The elevated FIGLU-to- creatinine ratios in AN patients given N2O returned toward baseline by postoperative day 1.

Discussion: These findings suggest a minimal disruptive effect of N2O on vitamin B12 and folate status in patients, even in elderly individuals who tend to be deficient in these vitamins and in those who receive N2O for prolonged surgical procedures.