Seminole Community College Paramedic 1 Medication Handout #4 Adenocard (Adenosine)

Description-

Adenosine is an endogenous nucleoside occurring in all cells of the body. Adenosine is not chemically related to other antiarrhythmic drugs. Adenocard® (adenosine injection) is a sterile, nonpyrogenic solution for rapid bolus intravenous injection.

Mechanism of Action

Adenocard (adenosine injection) slows conduction time through the A-V node, can interrupt the reentry pathways through the A-V node, and can restore normal sinus rhythm in patients with paroxysmal supraventricular tachycardia (PSVT), including PSVT associated with Wolff-Parkinson-White Syndrome. Adenocard is antagonized competitively by methylxanthines such as caffeine and theophylline, and potentiated by blockers of nucleoside transport such as dipyridamole. Adenocard is not blocked by atropine.

Hemodynamics

The intravenous bolus dose of 6 or 12 mg Adenocard (adenosine injection) usually has no systemic hemodynamic effects. When larger doses are given by infusion, adenosine decreases blood pressure by decreasing peripheral resistance.

Pharmacokinetics

Intravenously administered adenosine is rapidly cleared from the circulation via cellular uptake, primarily by erythrocytes and vascular endothelial cells. This process involves a specific transmembrane nucleoside carrier system that is reversible, nonconcentrative, and bidirectionally symmetrical. Intracellular adenosine is rapidly metabolized either via phosphorylation to adenosine monophosphate by adenosine kinase, or via deamination to inosine by adenosine deaminase in the cytosol. Since adenosine kinase has a lower K_m and V_{max} than adenosine deaminase, deamination plays a significant role only when cytosolic adenosine saturates the phosphorylation pathway. Inosine formed by deamination of adenosine can leave the cell intact or can be degraded to hypoxanthine, xanthine, and ultimately uric acid. Adenosine monophosphate formed by phosphorylation of adenosine is incorporated into the high-energy phosphate pool. While extracellular adenosine is primarily cleared by cellular uptake with a half-life of less than 10 seconds in whole blood, excessive amounts may be deaminated by an ecto-form of adenosine

deaminase. As Adenocard requires no hepatic or renal function for its activation or inactivation, hepatic and renal failure would not be expected to alter its effectiveness or tolerability.

INDICATIONS AND USAGE:

Intravenous Adenocard (adenosine injection) is indicated for the following:

Conversion to sinus rhythm of paroxysmal supraventricular tachycardia (PSVT), including that associated with accessory bypass tracts (Wolff-Parkinson-White Syndrome). When clinically advisable, appropriate vagal maneuvers (e.g., Valsalva maneuver), should be attempted prior to Adenocard administration.

It is important to be sure the Adenocard solution actually reaches the systemic circulation (see DOSAGE AND ADMINISTRATION).

Adenocard does not convert atrial flutter, atrial fibrillation, or ventricular tachycardia to normal sinus rhythm. In the presence of atrial flutter or atrial fibrillation, a transient modest slowing of ventricular response may occur immediately following Adenocard administration.

CONTRAINDICATIONS:

Intravenous Adenocard (adenosine injection) is contraindicated in:

- 1. Second- or third-degree A-V block (except in patients with a functioning artificial pacemaker).
- 2. Sinus node disease, such as sick sinus syndrome or symptomatic bradycardia (except in patients with a functioning artificial pacemaker).
- 3. Known hypersensitivity to adenosine.

WARNINGS:

Heart Block

Adenocard (adenosine injection) exerts its effect by decreasing conduction through the A-V node and may produce a short lasting first-, second- or third-degree heart block. Appropriate therapy should be instituted as needed. Patients who develop high-level block on one dose of Adenocard should not be given additional doses. Because of the very short half-life of adenosine, these effects are generally self-limiting.

Transient or prolonged episodes of asystole have been reported with fatal outcomes in some cases. Rarely, ventricular fibrillation has been reported following Adenocard administration, including both resuscitated and fatal events. In most instances, these cases were associated with the concomitant use of digoxin and, less frequently with digoxin and verapamil. Although no causal relationship or drug-drug interaction has been established, Adenocard should be used with

caution in patients receiving digoxin or digoxin and verapamil in combination. Appropriate resuscitative measures should be available.

Arrhythmias at Time of Conversion

At the time of conversion to normal sinus rhythm, a variety of new rhythms may appear on the electrocardiogram. They generally last only a few seconds without intervention, and may take the form of premature ventricular contractions, atrial premature contractions, sinus bradycardia, sinus tachycardia, skipped beats, and varying degrees of A-V nodal block. Such findings were seen in 55% of patients.

Bronchoconstriction

Adenocard (adenosine injection) is a respiratory stimulant (probably through activation of carotid body chemoreceptors) and intravenous administration in man has been shown to increase minute ventilation (Ve) and reduce arterial PCO₂ causing respiratory alkalosis.

Adenosine administered by inhalation has been reported to cause bronchoconstriction in asthmatic patients, presumably due to mast cell degranulation and histamine release. These effects have not been observed in normal subjects. Adenocard has been administered to a limited number of patients with asthma and mild to moderate exacerbation of their symptoms has been reported. Respiratory compromise has occurred during adenosine infusion in patients with obstructive pulmonary disease. Adenocard should be used with caution in patients with obstructive lung disease not associated with bronchoconstriction (e.g., emphysema, bronchitis, etc.) and should be avoided in patients with bronchoconstriction or bronchospasm (e.g., asthma). Adenocard should be discontinued in any patient who develops severe respiratory difficulties.

PRECAUTIONS:

Drug Interactions

Intravenous Adenocard (adenosine injection) has been effectively administered in the presence of other cardioactive drugs, such as quinidine, beta- adrenergic blocking agents, calcium channel blocking agents, and angiotensin converting enzyme inhibitors, without any change in the adverse reaction profile. Digoxin and verapamil use may be rarely associated with ventricular fibrillation when combined with Adenocard (see WARNINGS). Because of the potential for additive or synergistic depressant effects on the SA and AV nodes, however, Adenocard should be used with caution in the presence of these agents. The use of Adenocard in patients receiving digitalis may be rarely associated with ventricular fibrillation (see WARNINGS).

The effects of adenosine are antagonized by methylxanthines such as caffeine and theophylline. In the presence of these methylxanthines, larger doses of adenosine may be required or adenosine may not be effective. Adenosine effects are potentiated by dipyridamole. Thus, smaller doses of adenosine may be effective in the presence of dipyridamole. Carbamazepine has been reported to increase the degree of heart block produced by other agents. As the primary effect of adenosine is to decrease conduction through the A-V node, higher degrees of heart block may be produced in the presence of carbamazepine.

Pediatric Use

No controlled studies have been conducted in pediatric patients to establish the safety and efficacy of Adenocard for the conversion of paroxysmal supraventricular tachycardia (PSVT). However, intravenous adenosine has been used for the treatment of PSVT in neonates, infants, children and adolescents (see DOSAGE AND ADMINISTRATION).1

Geriatric Use

Clinical studies of Adenocard did not include sufficient numbers of subjects aged 65 and over to determine whether they respond differently from younger subjects. Other reported clinical experience has not identified differences in responses between elderly and younger patients. In general, Adenocard in geriatric patients should be used with caution since this population may have a diminished cardiac function, nodal dysfunction, concomitant diseases or drug therapy that may alter hemodynamic function and produce severe bradycardia or AV block.

ADVERSE REACTIONS:

The following reactions were reported with intravenous Adenocard (adenosine injection) used in controlled U.S. clinical trials. The placebo group had a less than 1% rate of all of these reactions.

Cardiovascular Facial flushing (18%), headache (2%), sweating, palpitations, chest pain, hypotension (less than 1%).

Respiratory Shortness of breath/dyspnea (12%), chest pressure (7%), hyperventilation, head pressure

(less than 1%).

Central
Nervous
System

Lightheadedness (2%), dizziness, tingling in arms, numbness (1%), apprehension, blurred vision, burning sensation, heaviness in arms, neck and back pain (less than 1%).

Gastrointestinal Nausea (3%), metallic taste, tightness in throat, pressure in groin (less than 1%). Also, in post-market clinical experience with Adenocard, cases of prolonged asystole, ventricular tachycardia, ventricular fibrillation, transient increase in blood pressure, bradycardia, atrial

fibrillation, and bronchospasm, in association with Adenocard use, have been reported (see WARNINGS).

OVERDOSAGE:

The half-life of Adenocard (adenosine injection) is less than 10 seconds. Thus, adverse effects are generally rapidly self-limiting. Treatment of any prolonged adverse effects should be individualized and be directed toward the specific effect. Methylxanthines, such as caffeine and theophylline, are competitive antagonists of adenosine.

DOSAGE AND ADMINISTRATION:

For rapid bolus intravenous use only.

Adenocard® (adenosine injection) should be given as a rapid bolus by the peripheral intravenous route. To be certain the solution reaches the systemic circulation, it should be administered either directly into a vein or, if given into an IV line, it should be given as close to the patient as possible and followed by a rapid saline flush.

Adult Patients

The dose recommendation is based on clinical studies with peripheral venous bolus dosing.

Central venous (CVP or other) administration of Adenocard has not been systematically studied.

The recommended intravenous doses for adults are as follows:

Initial dose: 6 mg given as a rapid intravenous bolus (administered over a 1-2 second period). **Repeat administration:** If the first dose does not result in elimination of the supraventricular tachycardia within 1-2 minutes, 12 mg should be given as a rapid intravenous bolus. This 12 mg dose may be repeated a second time if required.

Pediatric Patients

The dosages used in neonates, infants, children and adolescents were equivalent to those administered to adults on a weight basis.

Pediatric Patients with a Body Weight < 50 kg.

Initial dose: Give 0.05 to 0.1 mg/kg as a rapid IV bolus given either centrally or peripherally. A saline flush should follow.

Repeat administration: If conversion of PSVT does not occur within 1-2 minutes, additional bolus injections of adenosine can be administered at incrementally higher doses, increasing the amount given by 0.05 to 0.1 mg/kg. Follow each bolus with a saline flush. This process should continue until sinus rhythm is established or a maximum single dose of 0.3 mg/kg is used.

Pediatric Patients with a Body Weight > 50 kg:

Administer the adult dose.

Doses greater than 12 mg are not recommended for adult and pediatric patients.

NOTE: Parenteral drug products should be inspected visually for particulate matter and discoloration prior to administration.

HOW SUPPLIED:

Adenocard® (adenosine injection) is supplied as a sterile, non-pyrogenic solution in normal saline.

NDC 0469-8234-12 Product Code 823412 6 mg/2 mL (3 mg/mL) in a 2 mL (fill volume) *Ansyr®* plastic disposable syringe, in a package of ten.

NDC 0469-8234-14 Product Code 823414 12 mg/4 mL (3 mg/mL) in a 4 mL (fill volume) *Ansyr®* plastic disposable syringe, in a package of ten.

Store at controlled room temperature 15°- 30°C (59°- 86°F).

DO NOT REFRIGERATE as crystallization may occur. If crystallization has occurred, dissolve crystals by warming to room temperature. The solution must be clear at the time of use. Contains no preservatives. Discard unused portion.

Plastic syringes may require needle or blunt. To prevent needle-stick injuries, needles should not be recapped, purposely bent or broken by hand.

Rx Only