YR 2 PHARMACOLOGY UNIT EXAMINATION 2 -- MARCH 29, 1999. CHOOSE THE SINGLE <u>BEST</u> ANSWER FOR QUESTIONS 1 - 75.

- 1. Cyclosporine:
 - A. Metabolized to phosphoramide mustard.
 - B. Binds to CD3 receptor on lymphocytes.
 - C. Binds to cyclophilins and inhibits calcineurin.
 - D. Increased trapping of nuclear factor kB.
 - E. Blocks alpha chain of interleukin-2 receptor.

2. Tacrolimus:

- A. Inhibition of inosine monophosphate dehydrogenase.
- B. Interferes with signal transduction in T-lymphocytes during G1.
- C. Blocks activated interleukin-2 receptor.
- D. Metabolized to thioinosinic acid and thioguanylic acid.
- E. Inhibits calcineurin dependent translocation of NF-AT.
- 3. Which of the following is a property of glucocorticoids that accounts, in part, for their use as immunosuppressives?
 - A. Metabolized to phosphoramide mustard.
 - B. Binds to CD3 receptor on lymphocytes.
 - C. Binds to cyclophilins and inhibits calcineurin.
 - D. Increased trapping of nuclear factor kB.
 - E. Blocks alpha chain of interleukin-2 receptor.

- 4. Azathiaprine:
 - A. Inhibition of inosine monophosphate dehydrogenase.
 - B. Interferes with signal transduction in T-lymphocytes during G1.
 - C. Blocks activated interleukin-2 receptor.
 - D. Metabolized to thioinosinic acid and thioguanylic acid.
 - E. Inhibits calcineurin dependent translocation of NF-AT.
- 5. Which of the following agents has the greatest potential for inducing a hypoglycemic reaction:
 - A. Troglitazone.
 - B. Glyburide.
 - C. Dexamethasone
 - D. Metformin.
 - E. Acarbose
- 6. Which of the following insulin preparations would remain effective in maintaining near normal glucose levels 8 hrs post sub-cutaneous injection:
 - A. NPH insulin
 - B. Insulin lispro
 - C. Crystalline insulin
 - D. Porcine insulin
 - E. Regular insulin
- 7. Which agent is associated with the greatest risk of inducing lactic acidosis in a patient suffering from NIDDM:
 - A. Troglitazone.
 - B. Glyburide.
 - C. Dexamethasone.
 - D. Metformin.
 - E. Acarbose.

- 8. Causes of hypothyroidism include all of the following, EXCEPT:
 - A. Insufficiency of dietary iodine
 - B. Excess TIF release by pituitary
 - C. Presence of goitrogens (agents which inhibit iodine organification) in diet.
 - D. Autoimmune destruction of thyroid
 - E. Graves
- 9. The best immediate treatment for an acute episode of thyroid storm is:
 - A. Thyrotropin
 - B. Radioactive iodine
 - C. Dexamethasone.
 - D. Reverse T3.
 - E. Propanolol.
- 10. Which of the following would NOT be utilized in the treatment of Graves disease:
 - A. Sodium iodide
 - B. ¹³¹I
 - C. Reverse T_3
 - D. Propylthiouracil
 - E. Methimazole
- 11. An agent used to treat a severe migraine attack is:
 - A. Ergotamine.
 - B. Ergonovine.
 - C. Bromocryptine.
 - D. Propanolol.
 - E. Amitriptyline.

- 12. Synthetic ADH derivatives with antagonistic actions at V1 receptors, should:
 - A. Function as vasodilators.
 - B. Function as vasoconstrictors.
 - C. Function as a diuretic.
 - D. Function as an anti-diuretic.
 - E. Be effective for treating nephrogenic diabetes insipidus.
- 13. An agent likely to produce unwanted diuresis:
 - A. Lithium carbonate.
 - B. Desmopressin.
 - C. Lypressin.
 - D. Angiotensin II.
 - E. Aldosterone.
- 14. A Class 1 antiarrhythmic agent that prolongs action potential duration without exhibiting reverse use dependence inhibition of K^+ channels:
 - A. Procainamide
 - B. Lidocaine
 - C. Flecainide
 - D. Esmolol
 - E. Amiodarone

- 15. Potent Na⁺ channel blocker (Class 1C antiarrhythmic agent); useful for supraventricular arrhythmias, producing increased mortality with treatment of premature ventricular contractions (PVCs) after a myocardial infarction.
 - A. Procainamide
 - B. Lidocaine
 - C. Flecainide
 - D. Esmolol
 - E. Amiodarone
- 16. Antiarrhythmic used i.v. in patients following myocardial infarction; exhibits significant neurologic side effects and can cause convulsions if delivered too rapidly
 - A. Procainamide
 - B. Lidocaine
 - C. Flecainide
 - D. Esmolol
 - E. Amiodarone
- 17. Isoproterenol:
 - A. Decreases L-type Ca²⁺ channel current
 - B. Directly inhibits Na-K-ATPase
 - C. Leads to cross tolerance with nitrates
 - D. Decreases heart rate in part by vagal reflex
 - E. Decreases diastolic blood pressure

- 18. Which one of the statements is CORRECT?
 - A. Toxic concentrations of digitalis lead to reduced impulse conduction
 - B. Toxic concentrations of digoxin produce a different spectrum of cardiac arrhythmias compared to toxic concentrations of digitoxin
 - C. The dosage of digoxin should be increased with coadministration of quinidine
 - D. Digoxin has a longer half-life for elimination than digitoxin
 - E. Digoxin increases AV conduction by a parasympathomimetic mechanism
- 19. A patient with symptoms of congestive heart failure is given combination therapy of digitalis, ACE inhibitors, and diuretics. Which of the following statements is correct?
 - A. Systolic but not diastolic blood pressure will be reduced.
 - B. Serum potassium levels will rise
 - C. Myocardial remodeling will increase
 - D. Reduced preload will reduce cardiac output
 - E. Heart rate will be reduced
- 20. To decrease O_2 consumption in patients experiencing angina of effort:
 - A. Nitrates are used for venodilatation
 - B. Amyl nitrite is used prophylactically
 - C. Verapamil is contraindicated because of reflex sympathetic responses
 - D. Isosorbide dinitrate and nitroglycerin must be used alternatively to prevent tolerance
 - E. Dobutamine is used to increase contractility

- 21. Carvedilol:
 - A. Is an alpha-2 receptor antagonist with partial alpha-1 agonist activity
 - B. Must be used with caution in patients with congestive heart failure
 - C. Is a partial beta receptor agonist
 - D. Increases cardiac contractility
 - E. Is a beta-1 receptor agonist

22. Diltiazem:

- A. Is a negative inotrope
- B. Is unique among Ca²⁺-channel blockers in its high plasma protein binding
- C. Increases cardiac pacemaker rate
- D. Is more potent than dihydropyridines as a vasodilator
- E. Is a sodium channel blocker

23. Amiodarone:

- A. Is a Class 1A antiarrhythmic agent
- B. Is more likely than quinidine to prolong action potentials at lower heart rates
- C. Causes skin discolorations in a majority of patients
- D. Can cause pulmonary fibrosis
- E. Has negligible neurologic effects

24. Adenosine:

- A. Has a half life of 10 min
- B. Causes rapid depolarization of ventricular cells
- C. Inhibits AV conduction
- D. Inhibits SA node
- E. Must be delivered slowly to achieve its antiarrhythmic effect

- 25. Administration of digitalis prior to treatment with procainamide for atrial fibrillation is used for:
 - A. Enhancing antiarrhythmic effect by lowering extracellular ${\rm K}^{\scriptscriptstyle +}$
 - B. Lowering the risk of torsade de pointes
 - C. Offsetting the negative inotropy of procainamide
 - D. Offsetting the anti-muscarinic effects of procainamide
 - E. Increasing Na^{\dagger} channel binding
- 26. Nitrate therapy is useful in all EXCEPT which of the following:
 - A. Variant angina
 - B. Unstable angina
 - C. Reducing mortality after myocardial infarctions
 - D. Reducing cardiac output
 - E. Hypertension
- 27. Nifedipine therapy is associated with all EXCEPT which of the following:
 - A. Prolongation of Q-T interval
 - B. Decreased arterial tone in variant angina
 - C. Reflex sympathetic discharge
 - D. Half life of approximately 3 h
 - E. Decreased cardiac output in angina of effort

28. Propranolol:

- A. Increases cardiac contractility
- B. Reduces mortality in patients after myocardial infarction
- C. Generally exhibits very high bioavailability
- D. Is relatively specific for beta-1 receptors
- E. Increases rate of SA node phase 4 depolarization

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- 29. All of the following statements about the potential toxicity of chemicals are true **EXCEPT**:
 - A. Inhaled chemicals rapidly enter the circulation because of high pulmonary blood flow.
 - B. Highly lipophilic agents can produce dermal toxicity because of rapid absorption.
 - C. The kidneys are highly exposed to drugs because of high renal blood flow relative to tissue mass.
 - D. All chemicals possess a threshold dose below which no toxicity is observed.
 - E. The liver is highly exposed to i.v. administered drugs because of the first-pass effect.
- 30. Zero-order kinetics is observed in the elimination of drugs at tissue concentrations that are:
 - A. Below the Vmax for p450.
 - B. Sufficient to produce toxicity.
 - C. Saturating for metabolism and elimination.
 - D. Above 200 ug per ml.
 - E. Above 500 ug per ml.
- 31. Lead poisoning:
 - A. Can occur at blood lead concentrations <10
 - B. Can be diagnosed by observing increased concentrations of free erythrocyte porphyrin (FEP).
 - C. Has distinct symptoms that are not shared by other diseases.
 - D. Cannot be effectively treated with EDTA.
 - E. Is no longer a health problem because of the phase-out of leaded gasoline.

- 32. A patient is observed in the emergency room to exhibit the following symptoms: Severe headache, dizziness, dimness of vision, and throbbing in the temples. Oxygen-carrying capacity of red blood cells is significantly reduced. The following diagnosis is made:
 - A. Carbon monoxide poisoning with 10 to 20% saturation of blood.
 - B. Methanol poisoning.
 - C. Ethylene glycol poisoning.
 - D. Carbon monoxide poisoning with 30 to 40% saturation of blood.
 - E. Carbon monoxide poisoning with 20 to 30% saturation of blood.
- 33. Amyl nitrite is used as an antidote for which chemical?
 - A. Carbon monoxide.
 - B. Methanol
 - C. Cyanide
 - D. Organophosphates.
 - E. Ethylene glycol.
- 34. N-Acetylcysteine is used as an effective treatment of acetaminophen poisoning. The best explanation for its mechanism of protection is that N-acetylcysteine:
 - A. Repletes cellular glutathione content.
 - B. Supplies cells with cysteine for protein synthesis.
 - C. Acts as an antioxidant
 - D. Produces acetyl-CoA for the mitochondrial citric acid cycle
 - E. Binds the reactive metabolite of acetaminophen.

- 35. Which of the following statements about chelators is FALSE?
 - A. Dimercaprol: Some adverse cardiovascular effects, useful for As, Hg, Pb, but not for Cd.
 - B. 2,3-Dimercaptosuccinate (DMSA): Congener of BAL, more water soluble and less adverse effects than BAL because it acts primarily extracellularly.
 - C. Penicillamine: Used primarily for Cd and As, chronic use relatively safe with few adverse effects.
 - D. Desferoxamine: Binds iron very avidly, is toxic so used only when severity of poisoning justifies it.
 - E. EDTA: Efficient chelator of many divalent cations, penetrates membranes poorly, limited by binding essential calcium.
- 36. Which heavy metal is matched INCORRECTLY with its description?
 - A. Inorganic lead: Inhibits porphyrin metabolism; chronic poisoning causes symptoms including weakness, anorexia, tremor, GI disturbances.
 - B. Inorganic mercury: Binds to sulfhydryl groups; is corrosive and precipitates proteins.
 - C. Pentavalent arsenic: Uncouples cellular respiration.
 - D. Trivalent arsenic: Binds to sulfhydryl groups on proteins and enzymes.
 - E. Organic mercury: Binds to amino groups; accumulated primarily by liver.
- 37. Treatment of carbon monoxide poisoning involves all of the following **EXCEPT:**
 - A. Administration of amyl nitrite to react with hemoglobin to hasten CO excretion.
 - B. Removal of individual from source of exposure.
 - C. Maintenance of cardiovascular function in severely affected individuals.
 - D. Administration of pure oxygen.
 - E. Maintenance of respiration.

- 38. Correction of formate-induced acidosis is critical to a patient's survival in poisoning with which chemical?
 - A. Cyanide.
 - B. Ethylene glycol.
 - C. Methanol.
 - D. Organochlorine insecticides.
 - E. DDT.
- 39. Which vitamin is potentially the most toxic of all vitamins, is fat-soluble, and is treated by discontinuation of vitamin administration, reduction of intake of calcium, and generous administration of fluids?
 - A. Vitamin A.
 - B. Vitamin C.
 - C. Vitamin K.
 - D. Niacin.
 - E. Vitamin D.
- 40. Which statement about free radicals is FALSE?
 - A. Free radicals can be carbon-, nitrogen-, oxygen-, or sulfur-centered.
 - B. Free radicals are normally short-lived species that can be detected by chemical trapping and special methods.
 - C. Alterations in cellular macromolecules occur during the propagation phase.
 - D. Free radicals can be neutral, cationic, or anionic species.
 - E. All free radicals are toxic.

- 41. Which of the following compounds would not be given by oral administration due to problems with first pass metabolism?
 - A. Diethylstilbestrol
 - B. Ethinyl Estradiol
 - C. Norethindrone
 - D. Tamoxifen
 - E. Estradiol
- 42. Which of the following is a TRUE statement regarding Postmenopausal Hormone Therapy?
 - A. Estrogens are administered daily at a relatively large dose to induce normal, menstrual bleeding.
 - B. Estrogen replacement therapy is routinely prescribed as a prophylactic measure against osteoporosis.
 - C. In an attempt to mimic normal physiological conditions, it is recommended that estrogen be administered along with a small amount of androgen during hormone replacement therapy.
 - D. To lower the risk of endometrial cancer, hormone replacement therapy generally includes the combined administration of estrogens and progestins.
 - E. It is recommended that hormone replacement therapy be initiated with the largest dose consistent with the relief of symptoms.
- 43. Which of the following is NOT a therapeutic use of progestins?
 - A. Severe endometriosis
 - B. Refractory anemia
 - C. Postmenopausal hormone therapy
 - D. Contraception
 - E. Metastatic endometrial carcinoma

- 44. Which of the following is a FALSE statement regarding Mifepristone?
 - A. Mifepristone can act as both an antiprogestin and an antiglucocorticoid.
 - B. Mifepristone blocks ovulation by preventing the mid-cycle surge of gonadotropins.
 - C. Due to its antiglucorticoid effects, Mifepristone has been used as an anabolic agent.
 - D. Mifepristone is a derivative of norethindrone.
 - E. Mifepristone stimulates menstrual bleeding by stimulating the release of prostaglandins.
- 45. Which of the following is an adverse side-effect associated with the use of combination, steroidal oral contraceptive agents?
 - A. Cholestatic hepatitis, presenting as jaundice
 - B. Increased risk of endometrial carcinoma
 - C. Recurring nausea and vomiting
 - D. Hypertension
 - E. Increased risk of hepatic adenocarcinoma
- 46. Which of the following compounds is commonly used as an antiandrogen?
 - A. Flutamide
 - B. Fluoxymesterone
 - C. Oxandrolone
 - D. Mestranol
 - E. Clomiphene

- 47. In a patient already taking furosemide, digoxin and a potassium supplement, naproxen might
 - A. Lower serum potassium
 - B. Increase the glomerular filtration rate
 - C. Reduce inotropy
 - D. Reduce urine flow
 - E. Increase heart rate
- 48. Allopurinol decreases the metabolism of:
 - A. 5-fluorouracil
 - B. Probenicid
 - C. 6-mercaptopurine
 - D. Colchicine
 - E. Methotrexate
- 49. Cromolyn sodium:
 - A. Most effective when administered orally
 - B. Prevents the antigen-antibody reaction
 - C. Relaxes smooth muscle by its action on receptors
 - D. Administration prior to exercise or pollen exposure reduces asthmatic attacks
 - E. Inhibits cyclooxygenase 2

50. Zafirlukast:

- A. Long acting beta-agonist
- B. Inhibits 5'-lipoxygenase
- C. Direct acting bronchodilator
- D. Inhibits cyclooxygenase 2
- E. Blocks a leukotriene receptor

51. Aminophylline:

- A. Is a selective beta2-agonist
- B. May cause cardiac and CNS stimulation
- C. Activates adenyl cyclase
- D. Inhibits monoamine oxidase
- E. Inhibits xanthine oxidase
- 52. Salmeterol:
 - A. May induce skeletal muscle tremors in elderly patients
 - B. Inhibits phosphodiesterase
 - C. Is only effective as an aerosol
 - D. Has a shorter duration of action than terbutaline
 - E. Inhibits the antiinflammatory effects of leukotrienes
- 53. Which of the following drugs is least likely to raise serum renin levels?
 - A. Captopril
 - B. Propranolol
 - C. Nicardipine
 - D. Minoxidil
 - E. Hydralazine
- 54. Which of the following drugs would be the best choice to initiate treatment of mild hypertension in a patient who suffers from asthma:
 - A. Propranolol
 - B. Hydralazine
 - C. Labetalol
 - D. Captopril
 - E. Pindolol

- 55. Which of the following drugs is most likely to antagonize the antihypertensive effect of captopril:
 - A. Colchicine
 - B. Ibuprofen
 - C. Probenicid
 - D. Allopurinol
 - E. Propranolol
- 56. Which of the following drugs exerts its antihypertensive effects, in part, by blockade of alphal-receptors:
 - A. Clonidine
 - B. Pindolol
 - C. Labetalol
 - D. Propranolol
 - E. Metoprolol
- 57. Which of the following diuretics would be most appropriate in a patient who is already being treated for diabetes with glyburide and captopril for hypertension:
 - A. Furosemide
 - B. Hydrochlorothiazide
 - C. Amiloride
 - D. Spironolactone
 - E. Mannitol
- 58. Which of the following antihypertensive agents is most likely to cause reflex tachycardia:
 - A. Verapamil
 - B. Atenolol
 - C. Nifedipine
 - D. Diltiazem
 - E. Nicardipine

- 59. Losartan and captopril share all of the following properties EXCEPT:
 - A. Both reduce cardiac remodeling due to hypertension
 - B. Both lower peripheral resistance
 - C. Both increase bradykinin's half-life
 - D. Both reduce aldosterone secretion
 - E. Both increase renin levels
- 60. All of the following are true of propranolol EXCEPT:
 - A. Initially, it can cause a hypertension due to beta2receptor blockade
 - B. Oral dosages vary among individuals due to "first pass" metabolism
 - C. It increases renin and angiotensin II levels
 - D. It increases VLDL and decreases HDL
 - E. Upon abrupt withdrawal, it can cause severe tachycardia
- 61. Which of the following drugs is associated with a lupus-like syndrome:
 - A. Hydralazine
 - B. Diltazem
 - C. Verapamil
 - D. Losartan
 - E. Minoxadil
- 62. Which of the following drugs decreases the renal clearance of digoxin:
 - A. Hydralazine
 - B. Diltazem
 - C. Verapamil
 - D. Losartan
 - E. Minoxadil

- 63. In congestive heart failure, cardiac output is determined mainly by:
 - A. Diuretic dosage
 - B. Collagen matrix
 - C. Conductance
 - D. Compliance
 - E. Contractility
- 64. Which of the following is a compensatory mechanism in left ventricular dysfunction:
 - A. Reduce preload
 - B. Increase compliance
 - C. Increase atrioventricular conduction time
 - D. Reduce calcium influx
 - E. Increase norepinephrine
- 65. Increasing endothelin:
 - A. Increases bradykinin degradation
 - B. Causes pulmonary vasoconstriction
 - C. Increases renin turnover
 - D. Reduces norepinephrine
 - E. Increases myocardial compliance

- 66. Expression of the multidrug-resistance phenotype results in the appearance of an outward drug transport system which can prevent the accumulation of many anti-tumor agents and thereby antagonize drug toxicity. This phenomenon can be demonstrated
 - A. Only in drug-resistance malignant cells
 - B. Only in metastatic cancer cells
 - C. Only in cardiac muscle cells
 - D. Only in kidney tubules
 - E. In all rapidly-dividing cells
- 67. If a 5 ig/ml concentration of cytosine arabinoside will kill 40% of an S-phase tumor cell culture, a 10 ig/ml concentration is expected to kill what percentage of the total number of cells?
 - A. 80%
 - B. 64%
 - C. 60%
 - D. 50%
 - E. 45%
- 68. The principle advantage of the use of alkylating agents in chemotherapy is their ability to:
 - A. Spare the bone marrow from toxicity
 - B. Cross the blood-brain barrier
 - C. Kill both rapidly- and slowly-dividing tumor cells
 - D. Delay the development of drug resistance
 - E. Bind to plasma protein

- 69. Toxicity of bleomycin to the skin and lung is associated with:
 - A. Absence of bleomycin hydrolase in those tissues
 - B. Affinity of the drug for collagen
 - C. Sensitivity of rapidly-dividing cells to this drug
 - D. Absence of the multidrug resistance phenotype
 - E. Poor oxygenation thereby preventing radical formation
- 70. An important mode of drug resistance that results <u>from impaired</u> <u>signal-transduction pathways</u> in neoplastic cells is:
 - A. Enhanced outward drug transport
 - B. Lack of drug metabolism to pharmacologically-active products
 - C. Enhanced repair of DNA damage
 - D. Failure of cells to undergo apoptosis after DNA is damaged
 - E. Inhibition of drug-transport systems
- 71. Which of these adverse effects of chemotherapy is an example of drug-induced inhibition of DNA synthesis in normal cell types?
 - A. Skin and CNS toxicity of high-dose cytosine arabinoside
 - B. Cardiac toxicity of doxorubicin
 - C. Hemorrhagic cystitis after treatment with cyclophosphamide
 - D. Bone marrow depression induced by methotrexate
 - E. Pulmonary fibrosis caused by bleomycin

- 72. In patient with impaired liver function, the dose of doxorubicin must be reduced because the drug is:
 - A. Activated by liver enzymes
 - B. Detoxified by liver enzymes
 - C. Excreted via the liver and bile
 - D. Distributed via binding to lipoproteins
 - E. Is a powerful hepatotoxin