

## **Criteria necessary for accreditation of the subject “General pharmacology and toxicology”**

### **1) Mandatory classes:**

- Course pharmacology and toxicology, 6<sup>th</sup> semester, Wed. 9:15-10:45 am+ Fri. 10:15-11:45

### **2) Concordant class**

- Lecture pharmacology and toxicology, 5<sup>th</sup> and 6<sup>th</sup> semester, Tue.-Thurs. 8:15-9:00 am

### **3) Record of achievement:**

- Exam at the end of the 6<sup>th</sup> semester

### **4) Learning objective: pharmacology and toxicology**

## **1. General pharmacology and toxicology**

### **1.1 Pharmacodynamics**

- Drug-effects [(non-) receptor transmitted, reversible und irreversible]
- Signaling pathways for membrane and intracellular receptors
- Dose-effect curve (maximum effect, potency, receptor reserve)
- Agonists, partial agonists, inverse agonists
- Antagonists (reversible, irreversible, competitive, non-competitive)
- Plasma concentration-effect-correlation, Hysteresis
- Pharmacodynamic interactions
- Pharmacodynamic tolerance und sensibility
- Drug-dependency und –addiction

### **1.2 Pharmacokinetics**

- Resorption (mechanisms, significance of physicochemical characteristics)
- Distribution (mechanisms, significance of physicochemical characteristics)
- Elimination (systemic, pre-systemic)
  - Elimination by expulsion (renal, biliary, enteral etc.) - Elimination by metabolism (hepatic, enteral etc.)

Phase-I- und Phase-II-metabolism

Pharmacogenetics

- Pharmacokinetic variables
  - Systemic availability (“Bioavailability ”)
  - Elimination half-life, terminal half-life

- Plasma (Blood)-Clearance (total, renal, extra renal clearance) – Distribution compartments

- Culminating, „deep“ compartment
- Initial dose und maintenance dose
- Linear und non-linear Kinetic
- Relation between pharmaceutical effects, time, and effect duration
- Pharmacokinetic interactions
  
- Pharmacokinetic tolerance

### **1.3 General toxicology**

• Areas of toxicology

- Drug toxicology

- tissue toxicology

- food toxicology

- environmental toxicology

• Exposition and dose-effect-correlation

- acute, chronic

- local, systemic

- Significance of the intake path (metabolic activation/inactivation) - dose-effect-correlation

• Quality of toxic affects

- Acute toxicity

- Chronic toxicity

- Teratogenicity, Embryo toxicity

- Carcinogenicity

- Immunotoxicity

- Fertility problems

• Principles risk assessment

- Determining toxicity (acute, subacute, subchronic, chronic)

- Determining kinetics and biotransformation

- Mutagenicity examinations

- Carcinogenicity examination

- Tests for reproductive toxicity (Fertility, embryotic harm)

- evaluation concepts with determination of norm-intervals (ADI, MAK, BAT, TRK)

• Human biomonitoring (HBM)

- Exposure monitoring (internal dose); effect monitoring

- Foreign body spectrum; advantages and limits of HBM

- Matrices (blood, urine, saliva)

- evaluation (biologic limit values (BLV, human biomonitoring values I and II)
- risk assessment by utilizing the internal dose of physical properties
- cases

## **1.4 Drug-development**

- Development
- Approval and monitoring

## **1.5 Dogmatic treatment areas**

- Phytotherapy
- Homeopathy
- Anthroposophy therapy

## **2. Specific Pharmacology and toxicology**

(all pharmaceutical groups include the following aspects: typical active ingredients, mechanism of action, effects, Pharmacokinetics, usage and contraindications, side effects and interactions)

### **2.1 Pharmacology des autonomic nervous system**

#### *2.1.1 Sympathetic nervous system*

- Synthesis, storage, release and inactivation of Noradrenalin and Adrenalin
- α- und β-Adrenoceptors and their subtypes
- Receptor-distribution in the organism; synaptic localization
- Signaling pathways; sensibilisation und desensibilisation – transmitted effects
- Drugs with affects on the noradrenergic transmission by
  - Synthesis-inhibition and production of false transmitters (α-Methyldopa)
  - Inhibition of vesicular storage (Reserpine)
  - Inhibition or increase of transmitter release (Clonidine, Moxonidine; α<sub>2</sub>-adrenoceptor-agonist and -antagonists, Guanethidine, indirect sympathomimetics)
  - Inhibition of transmitter-inactivation (inhibitors of re-uptake; inhibitors of MAO-A, MAO-B or COMT)
- Drugs with effects on α-adrenoceptors
  - Subtype-selective and non-selective agonists (i.e. Phenylephrine, Oxymetazoline)
  - Subtype-selective and non-selective antagonists (i.e. Phentolamine, Prazosin)
  - partial agonists (i.e. "mothercorn" alkaloids)
- Drugs with effects on β-adrenoceptors
  - Subtype-selective and non-selective agonists (i.e. Isoprenaline, Fenoterol,

Salmeterol)

- Hybrid agonists with effects on  $\alpha$ - und  $\beta$ -adrenoceptors (i.e. Dobutamine)
- Subtype-selective and non-selective antagonists (i.e. Atenolol, Propranolol, Metoprolol, Bisoprolol, Esmolol)
- Hybrid antagonists with effects on  $\alpha$ - und  $\beta$ -Adrenoceptors or NO- release (i.e. Carvedilol, Nebivolol)
- partial agonists (i.e. Pindolol)

### *2.1.2 Parasympathetic nervous system and other cholinergic systems*

- Synthesis, storage, release and inactivation of Acetylcholine
- Cholinoceptors (muscarinic, nicotinic) and their subtypes
- Distribution in organisms, synaptic localizations
- Signaling pathways, sensitisation und desensibilisation –transmitted effects
- Drugs with effects on the cholinergic transmission by inhibition
- Transmitter-release (Botulinum-Neurotoxin; Loperamid)
- Transmitter-inactivation (Cholinesterase-Inhibitors)
- Drugs with affects on muscarinic cholinoceptors
- Agonists with tertiary or quaternary nitrogen (i.e. Pilocarpine, Carbachol)
- Antagonists tertiary or quaternary nitrogen (i.e. Atropine, Butylscopolamine)
- Drugs with effects on nicotinic cholinoceptors of the muscular type
- Agonists (depolarizing muscle relaxants: Suxamethonium)
- Antagonists (non- depolarizing muscle relaxants: i.e. Atracurium, Rocuronium, Pancuronium, historically: Tubocurarine)
- Sugammadex
- Drugs with effects on nicotinic cholinoceptors of the neuronal type
- Agonists on ganglion type nicotinic receptors (Nicotine)
- Antagonists on ganglion type nicotinic receptors (Ganglionic blockers)
- Addendum: myotropic muscle relaxants (Dantrolene)

## **2.2 Pharmacology des central nervous system**

### *2.2.1 Amino acid -Transmitters of the CNS*

- GABA (GABAergic neuron systems, GABA-synapsis, GABA-receptors)
- Glycine (Glycinergic neuron systems, Glycinergic-synapsis, Glycine-receptors)
- Glutamate (Glutamatergic neuron systems, Glutamate-Synapsis, Glutamate-receptors)

### *2.2.2 Narcotics*

- Inhalation narcotics (Gases i.e N<sub>2</sub>O, Xenon, vapours: i.e. Isoflurane, Sevoflurane, Desflurane)
- Injection narcotics (Barbiturates, Ketamine, Etomidate, Propofol, Benzodiazepine, Dexmedetomidine)

#### *2.2.3 Hypnotics/Anxiolytics (Tranquilizers)*

- Benzodiazepines (BZD-receptor on the GABAa-receptor chloride ion channel complex
  - Agonists for the BZD-receptor with short-and long-term effects
  - Antagonists for the BZD-receptor (Flumazenil)
- Zopiclone, Zolpidem, Zaleplon
- Chloral hydrate and Clomethiazole
- Anxiolytics and sedatives from different drug groups: Opipramol, Trimipramine
- Antihistamines with hypnotic effects (i.e. Promethazine, Diphenhydramine)
- Addendum: central muscle relaxants
  - Benzodiazepines (i.e. Tetrazepam)
  - GABA<sub>B</sub>-receptor agonists (Baclofen)

#### *2.2.4 Antiepileptics (Anticonvulsants)*

- Antiepileptics with effects on the GABAergic transmission
  - Substances with effects on the GABAa-receptor (BZD, Phenobarbital, Topiramate)
  - Substances with effects on GABA-inactivation (Valproate, Vigabatrin, Tiagabine)
  - Substances with effects on the GABA-releasing vesicle protein SV2A (Levetiracetam)
- Antiepileptics with Na<sup>+</sup> -channel-blocking effect (Carbamazepine, Phenytoin, Loamotrigine, Valproate)
- Antiepileptics with CA2+-channel-blocking effect (Pregabalin, Gabapentin, Ethosuximide, Topiramate, Valproate)
- Antiepileptics with effects in the glutamatergic transmission (i.e. Lamotrigine, Valproate)
- Substances with effects on Glutamate-receptors (i.e. Topiramate)

#### *2.2.5 Anti-Parkinson medication*

- Dopamine as a CNS-transmitter
  - Dopaminergic neuron systems, dopaminergic synapses
  - Dopamine-receptors, their signaling pathways and transmitted effects
- Levodopa (Combination with Dopa Decarboxylase-Inhibitors (i.e. Benserazide))
- Dopamine-receptor agonists (Pramipexole, Ropinirole, Apomorphine, historically ergoline-derivatives (i.e. Lisurid, Bromocriptine))
- COMT-inhibitors (Entacapone, Tolcapone)
- MAO-B inhibitors (Selegiline, Rasagiline)
- Muscarinic receptor- antagonists (i.e. Biperiden, Trihexyphenidyl, Metixen)
- NMDA antagonists (Amantadin, Budipin)

## *2.2.6 Antipsychotic medication*

- First-generation antipsychotics
  - Phenothiazines (i.e. Chlorpromazine, Levomepromazine, Perphenazine)
  - Thioxanthenes (i.e. Chlorprothixene, Flupentixol)
  - Butyrophenones (i.e. Haloperidol, Benperidol, Melperone, Pipamperon)
  - Diphenylbutylpiperidines (i.e. Fluspirilene, Pimozide)
- Second-generation antipsychotics (i.e. Clozapine, Olanzapine, Quetiapine, Risperidone, Aripiprazole)

## *2.2.7 Antidepressants*

- Mechanism of drug effect
- Non-selective monoamine-reuptake-inhibitors
  - Tricyclic and tetracyclic AD (Imipramine, Clomipramine, Nortriptyline, Amitriptyline, Doxepin, Maprotiline)
  - Venlafaxine, Duloxetine, Bupropion
- Selective Serotonin-reuptake-inhibitors (Fluoxetine, Citalopram, Paroxetine, Sertraline)
- Selective noradrenaline-reuptake-inhibitors (Reboxetine)
- a2-adrenoceptor-antagonists (Mirtazapine, Mianserin)
- Melatonin-receptor agonists (Agomelatine)
- MAO-inhibitors (Tranylcypromine, Moclobemide)
- St. John's Wort (Hyperforin)

## *2.2.8 Mood-stabilizers*

- Lithium carbonate
- Carbamazepine and Valproate

## *2.2.9 clinically used psychostimulants*

- Methylphenidate, Atomoxetine, Modafinil

## *2.2.10 Antidementives*

- Acetylcholinesterase- inhibitors (i.e. Rivastigmine, Donepezil)
- NMDA- antagonists (Memantine)

## *2.2.11 Substances with abuse and addiction potential*

- Substances with activating effects on the mesolimbic dopamine system
  - Psychostimulants Amphetamine and its derivatives, cocaine;
  - Opioids

-Nicotine

-clinically used psychostimulants: Methylphenidate, Atomoxetine, Modafinil

- Psychotomimetics: 5HT2a-receptor-stimulating substances

- direct-working substances: Indoleamine (i.e. LSD), Phenylethylamines (i.e. Mescaline)

- indirect-working substances (i.e. Ecstasy)

- Psychotomimetics: NMDA receptor blocking substances

- Phencyclidine, Ketamine

- Ethanol

- Substances with exponentiating effects on GABAa receptor- transmitted GABA effects

- Benzodiazepines, barbiturates

- Ethanol

- Adenosine receptor-blocking substances

- Methylxanthine (Caffeine, Theophylline, Theobromine)

- Substances with Cannabinoid receptor- stimulating effects

- Cannabinoid receptors (CB1,CB2) and their endogenous ligands

- Tetrahydrocannabinol (Marijuana, Hashish)

## **2.3 Pharmacology of histamine**

- Occurrence, synthesis, storage, release and inactivation of histamine

- Histamine receptors

- Subtypes and signaling pathways

- Transmitted effects

- Pharmaceuticals with effects on histamine release

- $\beta_2$ - adrenoceptor- agonists (i.e. Adrenaline, Fenoterol)

- Mast cell degranulation inhibitors (Cromoglycate, Nedocromil)

- Histamine receptor antagonists

- H1- receptor- antagonists (i.e. Diphenhydramine, Meclizine, Cetirizine, Loratadine)

- H2- receptor- antagonist (i.e. Cimetidine, Ranitidine)

## **2.4 Pharmacology of serotonin/5-hydroxytryptamine (5-HT)**

- Occurrence, synthesis, storage, release and inactivation of 5-HT

- 5-HT-receptors
  - Subtypes and signaling pathways
  - Transmitted effects
- Pharmaceuticals with effects on the inactivation and the release of 5-HT
  - Inhibitors of 5-HT reuptake (SSRI)
  - Inhibitors of MAO-A (Moclobemid)
  - 5-HT releasing substances (i.e. Amphetamine, Ecstasy)
- 5-HT receptor agonists
  - Buspirone, Urapidil (5-HT1a)
  - Triptane (5-HT1b/d)
  - LSD (5-HT2a)
  - Metoclopramide (5-HT4)
  - Ergotamine, dihydroergotamine (partial agonists for various 5-HT receptors)
- 5-HT receptor antagonists
  - Methysergide (5-HT2 and 5-HT1d)
  - Ketanserin, Risperidone (5-HT2a)
  - Ondansetron etc. (5-HT3)

## **2.5 Pharmacology of analgesics**

### *2.5.1 Opioid-Analgesics*

- Endogenous Opioids
- Opioid-receptors und their subtypes
- Signaling pathways
- Transmitted effects
- low potent- and highly-potent Opioid-Analgesics
- partial Opioid receptor agonists and mixed agonists-antagonists
- Opioid receptor antagonists
- Antitussives

### *2.5.2 Antipyretic Analgesics*

- Non-selective COX-Inhibitors
  - acidic antipyretic analgesics (non-steroidal anti-inflammatory drugs)
  - non-acidic antipyretic analgesics (Paracetamol, Metamizole)
- Selective COX-2-inhibitors (i.e. Etoricoxib, Celecoxib)

### *2.5.3 Other Analgesics*

- Flupirtine and Nefopam
- NMDA receptor antagonists (Ketamine, Dextromethorphan, Dextromethadone)
- α<sub>2</sub>-adrenoceptor agonists (i.e. Clonidine)

#### *2.5.4 Adjuvant analgesic treatment*

- Antidepressants (i.e. Amitriptyline, Venlafaxine)
- Antiepileptics (i.e. Carbamazepine, Phenytoin, Gabapentin)
- Glucocorticoids (i.e. Prednisolone, Dexamethasone)
- Bisphosphonates (i.e. Alendronate, Risedronate)

### **2.6 Pharmacology of the uric acid metabolism**

- Substance for acute gout treatment
  - Colchicine
  - non-steroidal antiphlogistics and Glucocorticoids
- Pharmaceuticals with effects on the plasmatic uric acid level
  - Uricostatics (Allopurinol)
  - Rasburicase
  - Uricosurics (Benzbromarone, Probenecid)

### **2.7 Pharmacology of Ion channels**

#### *2.7.1 Local anesthetics*

- Local anesthetics of the ester-type (i.e. Procaine, Tetracaine)
- Local anesthetics of the acid amide type (i.e. Lidocaine, Bupivacaine)

#### *2.7.2 Antiarrhythmics*

- Class 1 antiarrhythmics (Na<sup>+</sup>-channel blockers: i.e. Ajmaline, Flecainid, Propafenone, Chinidene, Lidocaine)
- Class 2 antiarrhythmics (β-adrenoceptor antagonists: i.e. Propranolol, Metoprolol)
- Class 3 antiarrhythmics (K<sup>+</sup>-channel blockers: i.e. Amiodarone, Dronedarone, Sotalol)
- Class 4 antiarrhythmics (Ca<sup>++</sup>-channel blockers: i.e. Verapamil)
- other substances (Adenosine, cardiac glycosides)
- If-channel blockers (Ivabradine)

#### *2.7.3 Pharmacology of K<sup>+</sup>-channels*

- K<sup>+</sup>-channel types and their physiologic significance
- K<sup>+</sup>-channel opener (i.e. Nicorandil, Diazoxide, Minoxidil)
- K<sup>+</sup>-channel blockers
  - class III antiarrhythmics

- Sulfonylureas and Meglitinides
- Ivabradin

#### *2.7.4 Pharmacology of Ca<sup>++</sup>-channels*

- Ca<sup>++</sup>-channel types and their physiological significance
- Ca<sup>++</sup>-channel blockers
  - L-type calcium channel blockers (Dihydropyridines, Phenylalkylamines, Benzothiazepines)
  - T- channel blockers (Ethosuximides)

### **2.8 Pharmacology of positive-inotropic drugs**

#### *2.8.1 Inhibitors of Na<sup>+</sup>, K<sup>+</sup>-ATPase (Digitalis-Glycosides)*

- Digoxin, Digitoxin

#### *2.8.2 Sympathomimetics*

- Dopamine, Dobutamine

#### *2.8.3 Inhibitors of type III-Phosphodiesterase*

- Milrinone, Enoximone

### **2.9 Pharmacology of vasodilating drugs**

#### *2.9.1 Nitro vasodilators*

- Glyceroltrinitrat, Isosorbiddinitrat, Isosorbide-5-Mononitrate
- Molsidomin

#### *2.9.2 Inhibitors of type V-Phosphodiesterase*

- Sildenafil, Vardenafil

#### *2.9.3 Inhibitors of Angiotensin II-synthesis and-effect*

- Renin antagonists (Aliskiren)
- ACE- inhibitors (i.e. Ramipril, Captopril, Enalapril)
- AT1-receptor antagonists (sartans)

#### *2.9.4 Vasodilators with effects on ion channels*

- Ca<sup>++</sup>-channel blocker (L-type calcium channel blocker)
- K<sup>+</sup>-channel opener (i.e. Minoxidil, Diazoxide)

#### *2.9.5 Hydralazine and Dihydralazine*

### **2.10 Pharmacology of substances influencing the respiratory system**

#### *2.10.1 Bronchodilators*

- b2 adrenoceptor- agonists

- Substances for inhalation (i.e. Terbutaline, Fenoterol, Salmeterol)
- Substances for systemic use (i.e. Clenbuterol, Reproterol)
- Muscarinic receptor antagonists
  - Substances for inhalation (Ipratropium, Tiotropium)
- Theophylline

#### *2.10.2 Substances with an anti-inflammatory effect*

- Inhalation Glucocorticoids (i.e. Budesonide, Fluticasone, Beclomethasone-DP)
- Inhibitors of mediator-release (Cromoglycate, Nedocromil)
- CysLT1 receptor antagonists (Montelukast)

#### *2.11.1 Benzothiadiazines*

- Thiazides (i.e. Hydrochlorothiazide, Bendroflumethiazide)
- Thiazide-Analogs i.e. Chlortalidone, Xipamide, Indapamide

#### *2.11.2 Loop diuretics*

- Loop diuretics type Furosemide (i.e. Furosemide, Torasemide)

#### *2.11.3 Potassium sparing diuretics*

- Aldosterone antagonists (Spironolactone, Eplerenone)
- Epithelial sodium channel blockers (Triamterene, Amiloride)

#### *2.11.4 Carbonic anhydrase inhibitors (i.e. Acetazolamide, Dorzolamide)*

#### *2.11.5 Osmotic diuretics (i.e. Mannitol, Sorbitol)*

### **2.12 Pharmacology of blood and the blood producing system**

#### *2.12.1 Plasma substitutes*

- Colloids Plasma substitute (Dextran, Hydroxyethyl starch, Gelofusine)
- Homologous plasma

#### *2.12.2 Substances affecting blood production*

- Iron complexes (i.e. Ferric carboxymaltose)
- Vitamins (Vitamin B12, Folic acid)
- Hematopoietic growth factors
  - Erythropoietin (i.e. Epoetin alfa)
  - Granulocyte colony-stimulating factor (G-CSF: Filgrastim, Lenograstim)
  - Granulocyte-monocyte colony-stimulating factor (GM-CSF: Molgramostim)

## **2.13 Pharmacology of blood coagulation**

### *2.13.1 Inhibitors of thrombocyte aggregation*

- Acetylsalicylic acid
- ADP-antagonists
  - Thienopyridine-derivatives (Prasugrel, Clopidogrel, Ticlopidine, Ticagrelor)
  - Ticagrelor
- Glycoprotein (GP) IIb/IIIa-receptor-antagonists
  - Monoclonal antibodies against GPIIb/IIIa-receptor (Abciximab)
  - synthetic GPIIb/IIIa-receptor-antagonists
    - Peptidic substances (Eptifibatid)
    - Non-peptidic substances (i.e. Tirofiban)

### *2.13.2 Anticoagulation*

- Direct anticoagulation effect
  - Heparin (unfractionated Heparin; low molecular weight heparin)
  - Heparinoids (i.e. Danaparoid)
  - sulfate pentasaccharide (Fondaparinux)
  - Thrombin inhibitors (Hirudin, Lepirudin, Desirudin)
  - oral factor II- and factor Xa-inhibitors (Dabigatran, Rivaroxaban, Apixaban)
- Indirect anticoagulation effect
  - Coumarin derivatives (Phenprocoumon, Warfarin)
  - Factor Xa and IIa-Inhibitors (i.e. Apixaban, Rivaroxaban, Dabigatran)

### *2.13.3 Fibrinolytics (Thrombolytics)*

- Direct fibrinolytics (Urokinase, Alteplase etc.)
- Indirect fibrinolytics (Streptokinase, Anistreplase)

### *2.13.4 Antifibrinolytics*

- Aprotinin,  $\omega$ -Amino carbonic acid (i.e.  $\epsilon$ -Aminocaproic acid, Tranexamic)

## **2.14 Pharmacology of the gastrointestinal tract**

### *2.14.1 Substances with prokinetic effect on stomach and intestine*

- Dopamine D2 receptor-antagonists (i.e. Metoclopramide, Domperidone, Alizapride)
- 5-HT4-receptor agonists (i.e. Cisapride, Metoclopramide)
- Motilin-receptor agonists (i.e. Erythromycin)

### *2.14.2 Ulcer treatment*

- Inhibitors of the H<sup>+</sup>,K<sup>+</sup>-ATPase (proton pump inhibitors: i.e. Omeprazole, Pantoprazole)
- Histamine H<sub>2</sub>-receptor antagonists (i.e. Ranitidine)
- Misoprostol
- Antacids (Magnesium- and Aluminum- complexes), Sucralfate
- M<sub>1</sub>-receptor antagonists (Pirenzepin)

#### *2.14.3 Anti-inflammatory treatment for chronic-inflammatory intestinal diseases*

- Sulfasalazine
- 5-Aminosalicylic acid

#### *2.14.4 Laxatives*

- Lubricants (i.e. Glycerin)
- Bulk-forming laxatives (i.e. Agar-Agar, Methylcellulose)
- Saline und hyperosmotic agents (i.e. sodium phosphate, lactulose)
- Stimulant agents (i.e. Anthraquinone, Bisacodyl)

#### *2.14.5 Antidiarrheal Substances*

- Carbo medicinalis
- Loperamide, Diphenoxylate

#### *2.14.6 Antiemetics*

- Dopamine receptor antagonists (i.e. Metoclopramide, Domperidone)
- H<sub>1</sub>-receptor antagonists (i.e. Meclizine)
- Muscarinic receptor antagonists (Scopolamine)
- 5-HT<sub>3</sub>-receptor antagonists (i.e. Ondansetron)

### **2.15 Pharmacology of lipometabolism**

#### *2.15.1 Inhibitors of Hydroxymethylglutaryl-CoA-Reductase*

- Statins

#### *2.15.2 Fibrates*

- Clofibrate and Clofibrate derivatives (i.e. Etofibrate)
- Clofibrate-analogs (i.e. Gemfibrozil)

#### *2.15.3 Nicotinacid*

- Nicotinic acid
- Nicotinic acid-Analogs

#### *2.15.4 Bile acid sequestrant*

- Cholestyramine and Colestipol

#### *2.15.5 Cholesterol absorption inhibitor*

- $\beta$ -Sitosterol
- Ezetimibe

### **2.16 Pharmacology of diabetic metabolism malfunctions**

#### *2.16.1 Insulin*

- Human insulin (Normalinsulin)
- Intermediate-acting (i.e. NPH-Insulin)
- Human insulin analogues (i.e. Insulin lispro, Insulin glargin)

#### *2.16.2 Oral Anti-diabetics*

- Substances that enhance Insulin-release
  - Sulfonylureas (i.e. Glibenclamid, Glimepiride)
  - Meglitinide (i.e. Repaglinid)
- Substances that enhance receptor sensitivity
  - Metformin
  - Thiazolidindiones (Glitazone)
- Inhibitors of glucose absorption ( $\alpha$ -glucosidase inhibitors: i.e. Acarbose)
- Incretin (i.e. Exenatide, Sitagliptin)
- SGLT2-inhibitors (Dapagliflozin)

### **2.17 Pharmacology of hormonal systems**

#### *2.17.1 Thyroid pharmacology*

- Thyroid hormones (Levothyroxine, Liothyronine)
- Iodine
- Thyreostatics
  - Thiourea-Derivatives
  - Perchlorate
  - Radioiodine

#### *2.17.2 Adrenal cortex pharmacology*

- Glucocorticoids (Cortisol und synthetic Glucocorticoids)
- Mineralocorticoids (Aldosterone, Fludrocortisone)

#### *2.17.3 Gonadal pharmacology*

- Androgens (Testosterone, Testosterone undecanoate)
- Anti-androgens substances
- Steroids (Cyproteronacetat)
  - Non-Steroids (Bicalutamide, Enzalutamide)
  - 5 $\alpha$ -Reductase-inhibitors (Finasterid, Dutasteride)
  - Cyp17A1-inhibitor (Abiraterone)
- Estrogens (17 $\beta$ -Estradiol and its Derivatives: i.e. Ethinyl Estradiol)
- Anti-estrogenic substances
  - Clomiphene
  - Selective Estrogen receptor modulators (SERM: Tamoxifen, Raloxifene)
  - Aromatase-inhibitors (non-steroidal and steroid substances)
- Gestagens
  - Progesterone and Progesterone-derivatives (i.e. Medroxyprogesterone)
  - Nortestosterone-derivatives (i.e. Levonorgestrel, Gestodene)
- Hormonal contraception
  - Estrogen-Gestagen-Combinations – Mini pill, pure Gestagen drugs
  - Postcoital Contraception

## **2.18 Pharmacology of bone metabolism**

### *2.18.1 Calcium*

### *2.18.2 Vitamin D and its derivatives*

### *2.18.3 Substances with inhibitory effect on bone resorption*

- Bisphosphonate (i.e. Alendronate, Risedronate, Zoledronate)
- Estrogen und Raloxifen

### *2.18.4 Substances with stimulating effect on bone resorption*

- Teriparatid

## **2.19 Antibacterial pharmacology**

### *2.19.1 Inhibitors of cell wall synthesis*

- $\beta$ -Lactam-Antibiotics +  $\beta$ -Lactamase-Inhibitors
- Glycopeptides

### *2.19.2 Proteinsynthesis inhibitors*

- Aminoglycosides
- Tetracyclines
- Macrolids and Ketolids

- Lincosamids

#### *2.19.3 Chemotherapy affecting nucleoid acids*

- Fluorchinolones (Gyrase-inhibitor: 4 classifications according to the Paul-Ehrlich Gesellschaft)
- Nitroimidazole (i.e. Metronidazole)

#### *2.19.4 Folicacid antagonists*

#### *2.19.5 Anti-tuberculotics*

### **2.20 Antimycotic pharmacology**

#### *2.20.1 Antimycotics affecting structure and stability of cytoplasma membranes*

- Polyenes
- Imidazoles and Triazoles
- Allylamines

#### *2.20.2 Hemmstoffe der Zellwand-Biosynthese*

- Echinocandines

#### *2.20.3 Inhibitoren der Nucleinsäuresynthese*

- Flucytosin

### **2.21 Antiviral pharmacology**

#### *2.21.1 Virustatics for Herpes treatment (i.e. Aciclovir, Famciclovir)*

#### *2.21.2 Virustatics for CMV treatment (i.e. Ganciclovir, Cidofovir)*

#### *2.21.3 Virustatics for HIV treatment*

- Reverse transcriptase inhibitors
  - Nukleoside-analogues (NRTI)
  - Nukleotide-analogues (NTRTI)
  - Non-nucleoside reverse transcriptase-inhibitors (NNRTI)
  - Integrase-inhibitors
  - Protease-inhibitors
  - Entry-inhibitors
    - Co-receptor antagonists
    - Fusions-inhibitors
  - Current HIV treatment plans

#### *2.21.4 Antiviral treatment for Influenza*

- Amantadine
- Neuraminidase-inhibitors

#### *2.21.5 Antivirals for treatment of Hepatitis B and C*

- Nucleoside-analogues
- Ribavirin + Interferon Protease-inhibitors

### **2.22 Pharmacology of substances affecting protozoas and worms**

#### *2.22.1 Chemotherapy for protozoan infections*

- Malaria remedies
- Substances against toxoplasmosis
- Amoebiasis remedies

#### *2.22.2 Chemotherapy for helminthiasis*

- Substances that interrupt neuromuscular activity
- Other Anthelmintics

### **2.23 Pharmacology of substances affecting malignant tumors**

#### *2.23.1 Antimetabolites*

- Folic acid analogues (Methotrexate)
- Purine analogues (i.e. 6-Mercaptopurin)
- Pyrimidine analogues (i.e. Fluorouracil)

#### *2.23.2 Alkylating cytostatics*

- Nitrogenlost- derivatives (i.e. Cyclophosphamid)
- N-Nitrosourea- derivatives (i.e. Carmustin, Lomustin)
- Platin-Complexes (i.e. Cisplatin)

#### *2.23.3 Topoisomerase inhibitors*

- Topoisomerase I inhibitors (Topotecane)
- Topoisomerase II inhibitors (i.e. Etoposid)

#### *2.23.4 Mikrotubuli inhibitors*

- Vinca-Alkaloids (i.e. Vincristine)
- Taxanes (i.e. Paclitaxel)

### *2.23.5 Zytostatic antibiotics*

- Actinomycine (Dactinomycine)
- Anthracycline (i.e. Daunorubicine, Doxorubicine)
- Mitoxantron and Amsacrine
- Bleomycine
- Mitomycine

### *2.23.6 Hormones and Hormone antagonists*

- Estrogens
- Gestagens
- Gonadotropin-releasing hormone (GnRH)
- Tamoxifene
- Aromatase inhibitors

### *2.23.7 New treatment methods*

- Monoclonal antibodies (i.e. Rituximab, Trastuzumab)
- Immune system mediators (i.e. Interleukin 2, Interferon)
- Tyrosine-Kinase inhibitors (i.e. Imatinib)

## **2.24 Pharmacology of Substances affecting the immune system**

### *2.24.1 Fundamentals of humoral and cellular immunity*

### *2.24.2 Immunosuppressants*

- Cytotoxic immunosuppressants
  - Cyclophosphamide
  - Azathioprine
  - Methotrexate
  - Mycophenolate
- Activation inhibiting immunosuppressants
  - Glucocorticoids
  - Cyclosporine and Tacrolimus
  - Sirolimus
- Immunologic immunosuppressants
  - Monoclonal antibodies
  - Polyclonal antilymphocyte-Globulins

### *2.24.3 Immune system mediators and their antagonists*

- Immunoglobulin (passive immunization)
- Cytokines (Interleukins, Interferons)
- Antagonists of mediators or their receptor
  - TNF $\alpha$ -antagonists (Infliximab, Adalimumab, Etanercept)
  - Antagonists of the IL-1-receptor (Anakinra)
  - monoclonal antibodies of IL-2-receptors (Basiliximab, Daclizumab)

#### *2.24.4 Anti-rheumatic basic therapeutics*

- Methotrexate
- Sulfasalazine
- Hydroxychloroquine
- Leflunomide
- TNF $\alpha$ -antagonists (Infliximab, Adalimumab and Etanercept)

### **2.25 Particular Toxicology**

#### *2.25.1 Toxicology of chemical carcinogens*

- Chemical carcinogenicity
- Multiple step concept of carcinogenicity
- genotoxic carcinogenics, mechanism of action
- non-genotoxic carcinogenics, mechanism of action
- Carcinogenic substances
  - polycyclic aromatic carbohydrates
  - aromatic amines and heterocyclic amines
  - Nitro-complexes
  - halogenated Carbohydrates
  - other genotoxic complexes
  - Metals
  - Particles and fibers
  - other non-genotoxic carcinogenics

#### *2.25.2 Inhalation toxins and Methemoglobin builders*

- Inhalation toxins with systemic effects
  - Carbon-monoxide
  - Carbon-dioxide
  - Hydrogen cyanide, Antidote (=Methemoglobin builders) and their toxicology
  - Hydrogen sulfide
- Irritant gases
  - Development of toxic Lung edemas
  - Ozone
  - Sulfur dioxide

- Nitrous gases
- Chlorine

#### *2.25.3 Toxicology of organic solvents*

- Alcohols
  - Ethanol
  - Methanol
  - Glycols
- Aliphatic compounds
  - Halocarbon Aliphates (i.e. Chloroform, Dichloromethane)
  - Alkanes
- Aromatic compounds
  - Benzyl
  - Toluene

#### *2.25.4 Toxicology of pesticides*

- Insecticides
  - chlorinated hydrocarbon compounds
  - Cholinesterase-inhibitors
  - Pyrethroids
- Herbicides and Fungicides
  - Bispyridinium-derivatives
  - Pentachlorophenol
- Rodenticides
  - Anticoagulants
  - Thallium sulfates

#### *2.25.5 Toxicology of metal complexes*

- Lead compounds
  - inorganic lead silicate
  - organic lead compounds
- Mercury compounds
  - inorganic mercury compounds
  - organic mercury compounds
- Other metal complexes

- Arson compounds
- Cadmium complexes
- Thallium complexes
- Chelating agents as antidotes for metal intoxications
- EDTA
- DMPS
- D-Penicillamine
- Deferoxamine
- Prussian blue

#### *2.25.6 Natural substance toxicology*

- Bacterial Toxins
- Tetrodotoxin
- Botulinum-Toxins
- Tetanus-Toxins
- Mushroom toxins
- Muscarine
- Psilocybin
- Amanitins
- Phalloidin
- Aflatoxin, Ochratoxin
- Plant toxins
- Alkaloids
- Cardiac glycosides
- Furanocoumarin
- Animal toxins
- Bee- and wasp toxins
- Spider toxins
- Scorpion toxins
- Snake toxins

#### *2.25.7 Other areas of toxicology*

- Toxicology of tobacco consumption
- Nicotine-transmitted effects
- tobacco -specific nitrosamines
- other carcinogens in tobacco(smoke)
- Polyhalogenated aromatic carbohydrates
- Polyhalogenated dibenzo-p-dioxin
- Polyhalogenated dibenzofuran
- Polyhalogenated biphenyl

- Xenoestrogens and Phytoestrogens
  - Bisphenol A
  - Genistein
- Chemical hypersensitivity (MCS-Syndrome)

