

## Literaturverzeichnis

- 1 Apro MS. Controlling emesis related to cancer therapy. *Europ J Cancer* 1991 ;27:356–61 ( Review )
- 2 Apro MS. 5-HT<sub>3</sub>- receptor antagonists. An overview of their present status and future potential in cancer therapy-induced emesis. *Drugs* 1995;42: 551–68
- 3 Apro MS, Sessa C, Thurlimann B. SAKK 90/95: A randomised double-blind trial to compare the clinical efficacy of granisetron to metoclopramide, both combined to dexamethasone in the prophylaxis of chemotherapy-induced delayed emesis. *Proc Ann Soc Clin Oncol* 2000; 19: 600a
- 4 Agundez JAG, Ledesma MC, Ladero JM, Benitez J. Prevalence of CYP2D6 gene duplication and its repercussion on the oxidative phenotype in a white population. *Clin Pharmacol Ther* 1995; 57:265-69
- 5 Aklillu E, Persson I, Bertilsson L, Johansson I, Rodrigues F, Ingelman-Sundberg M. Frequent distribution of ultrarapid metabolizers of debrisoquine in an Ethiopian population carrying duplicated and multiduplicated functional CYP2D6 alleles. *J Pharmacol Exp Ther* 1996; 278: 441–46
- 6 Andrews PL, Hawthorn J. The neurophysiology of vomiting. *Baillieres Clinical Gastroenterology* 1988; 2:(1) 141–68
- 7 Andrews PLR, Davidson HIM. Activation of vagal afferent terminals by 5-hydroxytryptamine is mediated by the 5-HT<sub>3</sub>-receptor in the anaesthetized ferret. *J Physiol* 1990; 422: 92 P
- 8 Andrews PLR , Davis CJ. The mechanism of emesis induced by anti-cancer therapy. In: Andrews PLR , Sanger GJ eds. *Emesis in anti-cancer therapy*. London: Chapman and Hall 1993: 113–61
- 9 Andrews PLR , Rapeport WG , Sanger GJ. Neuropharmacology of emesis induced by anti-cancer therapy . *Trends Pharmacol Sci* 1988; 9: 334–41
- 10 Andrykowski MA. Definitial issues in the study of anticipatory nausea in cancer chemotherapy. *J Behav Med* 1986; 9: 33–41
- 11 Andrykowski MA, Jacobsen PB, Marks E et al. Prevalence, predictors and course of anticipatory nausea in women receiving adjuvant chemotherapy for breast cancer . *Cancer* 1988; 62: 2607–613

- 12 Antiemetic subcommittee of the Multinational Association of Supportive Care in Cancer: Prevention of chemotherapy- and radiotherapy-induced emesis: results of the Perugia Consensus Conference. *Ann Oncol* 1998; 9: 811–19
- 13 Aynacioglu AS, Sachse C, Brockmüller J et al. Allele frequencies of CYP2D6 and CYP2C19 in a Turkish population. Poster. *Microsomes, Drug Oxidation and Clinical Pharmacology*, Berlin, Sept. 1997: 20–21
- 14 Aynacioglu AS; Sachse C; Bozkurt A et al. Low frequency of defective alleles of cytochrome P450 enzymes 2C19 and 2D6 in the Turkish population. *Clin Pharmacol* 1999; 66: 185–92
- 15 Bathum L, Johansson I, Ingelman-Sundberg M, Horder M, Brosten K. Ultrarapid metabolism of sparteine : frequency of alleles with duplicated CYP2D6 genes in a Danish population as determined by restriction fragment length polymorphism and long polymerase chain reaction. *Pharmacogenetics* 1998; 8: 119–23
- 16 Berger AM, Clark–Snow RA. Nausea and vomiting. In : De Vita VT , Hellman S Rosenberg SA eds.: *Cancer : Principles and Practice of Oncology* . Fifth Edition. Philadelphia Lippincott-Raven Publishers 1997: 2705–714
- 17 Bernal ML, Sinnes B, Johansson I, Mc Lellan R, Dahl ML, Ingelman-Sundberg M. High frequency of subjects with multiple CYP2D6 genes in a North-Spanish population. *Eur J Clin Pharmacol* 1997 ; 52 (Suppl) :A160
- 18 Bertilsson L. Geographical/interracial differences in polymorphic drug oxidation. Current state knowledge of Cytochromes P450(CYP)2D6 and 2C19. *Clin Pharmacokinet* 1995;29 (3): 192-209
- 19 Bhandari P, Andrews PL. Preliminary evidence for the involvement of the putative 5-HT<sub>4</sub>-receptor in zacopride- and copper sulphate-induced vomiting in the ferret. *Eur J Pharmacol* 1991 ; 204: 273–80
- 20 Bleiberg H. A new class of antiemetics : the NK-1 receptor antagonists. *Curr Opin Oncol* 2000; 12 (4): 284–88
- 21 Bleiberg H, Van Belle S, Paridaens R, De Wasch G, Dirix LY, Tjens M . Compassionate use of a 5-HT<sub>3</sub>-receptor antagonist, tropisetron, in patients refractory to standard antiemetic treatment. *Drugs* 1992; 43 (Suppl 3): 27–32

- 22 Bloomer JC, Baldwin SJ, Smith GJ, Ayrton AD, Clarke SE, Chenery RJ. . Characterization of the cytochrome P450 enzymes involved in the in vitro metabolism of granisetron. *Br J Pharmacol* 1994; 38: 557-66
- 23 Blower PR. 5-HT<sub>3</sub> receptor antagonists and the cytochrome P450 system – clinical implications . *Cancer* 2000; 8: 405-14
- 24 Blower PR, Aapro M. Granisetron vs ondansetron : is it a question of duration of 5-HT<sub>3</sub> receptor blockade ? *Br J Cancer* 2002; 86: 1662–663
- 25 Brockmüller J, Kirchheiner J, Schmider J, Walter S, Sachse C, Müller-Oerlinghausen B, Roots I. The impact of the CYP2D6 polymorphism on haloperidol pharmacokinetics and on the outcome of haloperidol treatment. *Clin Pharmacol Ther* 2002; 4: 438–52
- 26 Brosen K, Gram LF, Haghfelt T, Bertilsson L. Extensive metabolizers of debrisoquine become poor metabolizer during quinidine treatment. *Pharmacol Toxicol* 1987; 60: 312–14
- 27 Brunsch U, Drechsler S, Eggert J, Grosse H, Ukena D, Imhoff W, Färber L. Prevention of chemotherapy-induced nausea and vomiting by tropisetron (Navoban) alone or in combination with other antiemetic agents. *Seminars Oncology* 1994; 5 (Suppl 9): 7–11
- 28 Brunsch U , Rufenacht E, Parker J, Drechsler S, De Bruijn K. Tropisetron in the prevention of chemotherapy-induced nausea and vomiting in patients responding poorly to previous conventional antiemetic therapy. *Ann Oncol* 1993; 4: 25–29
- 29 Casper J, Schmoll H. Therapie von chemotherapiebedingter Übelkeit und Erbrechen. *Arzt und Krankenhaus* 1994; 10: 313–25
- 30 Clark RA, Gralla RJ. Delayed emesis: a dilemma in antiemetic control . *Support Care Cancer* 1993; 1: 182–85
- 31 Cooper S, Georgiu V. The impact of cytotoxic chemotherapy – perspectives from patients, specialists and nurses. *Eur J Cancer* 1992; 28 A (Suppl.1): 536–38
- 32 Corrigan BW, Nicholls B, Thakrai B. Heterogeneity in systemic availability of ondansetron and granisetron following oral administration. *Drug Metab Dispos* 1999; 27: 110–12
- 33 Cubeddu LX. Mechanisms by which cancer chemotherapeutic drugs induce emesis. *Seminars in Oncology* 1992; 19 (Suppl.15): 2–13

- 34 Cubeddu LX, Hoffmann IS, Fuenmayor NT, Finn AL. Efficacy of ondansetron (GR 38032 F) and the role of serotonin in cisplatin-induced nausea and vomiting. *N Engl J Med* 1990; 322: 810–16
- 35 Cupissol DR, Serrou B, Canbel M. The efficacy of granisetron as a prophylactic anti-emetic and intervention agent in high-dose cisplatin-induced emesis. *Eur J Cancer* 1990; 26 (Suppl 1) , S23–S27
- 36 Dahl ML, Johansson I, Bertilsson L, Ingelman-Sundberg M, Sjöqvist F. Ultrarapid hydroxylation of debrisoquine in a Swedish population. Analysis of the molecular genetic basis. *J Pharmacol Exp Ther* 1995; 274: 516–20
- 37 Dahl ML, Johansson I, Palmertz MP, Ingelman-Sundberg M, Sjöqvist F. Analysis of the CYP2D6 gene in relation to debrisoquin and desipramine hydroxylation in a Swedish population. *Clin Pharmacol Ther* 1992; 51 (1): 12–17
- 38 Daly AK, Brockmöller J, Broly F et al. Nomenclature for human CYP2D6 alleles *Pharmacogenetics* 1996; 6: 193–201
- 39 Daly AK, Fairbrother KS, Andreassen OA, London SJ, Idle JR, Steen VM. Characterization and PCR-based detection of two different hybrid CYP2D7P/CYP2D6 alleles associated with the poor metabolizer phenotype. *Pharmacogenetics* 1996; 6: 319–28
- 40 Daly AK, Leathart JB, London SJ, Idle JR. An inactive cytochrome P450 (CYP2D6) allele containing a deletion and a base substitution. *Hum Genet* 1995; 95 (3): 337–41
- 41 De Boer-Dennert M, de Wit R, Schmitz PI, Djontono J, v Beurden V, Stoter G, Verweij J. Patient perceptions of the side-effects of chemotherapy: the influence of 5-HT<sub>3</sub>-Antagonists. *Brit J Cancer* 1997; 76 (8): 1055– 61
- 42 De Bruijn KM. The development of tropisetron in its clinical perspective. *Ann Oncol* 1993; 4: 19–23
- 43 Del Favero A, Roila F, Tonato M. Reducing chemotherapy-induced nausea and vomiting. Current perspectives and future possibilities. *Drug Safety* 1993; 9: 410–28
- 44 De Mulder PHM, Seynave C, Vermorken JB, van Liessum PA, Mols-Jevdevic S, Allman EL. Ondansetron compared with high-dose metoclopramide in prophylaxis of acute and delayed cisplatin-induced nausea and vomiting. *Ann Intern Med* 1990; 113: 834-40

- 45 Dixon CM, Colthup PV, Serabit-Singh CJ, Kerr BM, Boehlert CC, Park GR, Tarbit MH. Multiple forms of cytochrome P450 are involved in the metabolism of ondansetron in humans. *Drug Metab Dispos* 1995; 23: 1225-30
- 46 Dogliotti L, Antonacci RA, Paze E, Ortega C, Berruti A, Faggiulolo R. Three years experience with tropisetron in the control of nausea and vomiting in cisplatin-treated patients. *Drugs* 1992; 43 (Suppl.3): 6-10
- 47 Drechsler S, Bauer R. 5-HT<sub>3</sub>-Rezeptorantagonisten. *Arzneimitteltherapie* 1995; 13: 224–35
- 48 Eichelbaum M, Mineshita S, Ohnhaus EE. The influence of enzyme induction on polymorphic sparteine oxidation. *Brit J Clin Pharmacol* 1986; 22: 49-53
- 49 Evans WE, Relling MV. Concordance of P450 2D6 (debrisoquine hydroxylase) phenotype and genotype: inability of dextromethorphan metabolic ratios to discriminate reliably heterozygous and homozygous extensive metabolizers. *Pharmacogenetics* 1991; 1 (3):143-48
- 50 Fauser AA, Duclos B, Chemaissani A. Therapeutic equivalence of single oral doses of dolasetron mesilate and multiple doses of ondansetron for the prevention of emesis after moderately emetogenic chemotherapy. *Eur J Cancer* 1996; 32A: 1523-29
- 51 Fauser AA, Fellhauer M, Hoffmann M. Guidelines for antiemetic therapy: Acute emesis. *Europ J Cancer* 1999; 35: 361-70
- 52 Fiore JJ, Gralla RJ. Pharmacological treatment of chemotherapy-induced nausea and vomiting. *Cancer Invest* 1984; 2: 351-61
- 53 Firkusny L, Krömer HK, Eichelbaum M. In vitro characterization of cytochrome P450 catalyzed metabolism of the antiemetic tropisetron. *Biochem Pharmacol* 1995; 49: 1777-84
- 54 Fischer V, Baldeck JP, Tse FL. Pharmacokinetics and metabolism of the 5-hydroxytryptamine antagonist tropisetron after single oral doses in humans. *Drug Metab Dispos* 1992; 20: 603-07
- 55 Fischer V, Vickers AE, Heitz F, Mahadevan S, Baldeck JP, Minery P, Tynes R. The polymorphic cytochrome P-4502D6 is involved in the metabolism of both 5-hydroxytryptamin-antagonists, tropisetron and ondansetron. *Drug Metab Dispos* 1994; 22: 269-74

- 56 Fletcher CV, Acosta EP, Strykowski JM. Gender differences in human pharmacokinetics and pharmacodynamics. *J Adolesc Health* 1994; 15: 619-29
- 57 Fredrikson M, Husti T, Steineck G, Fürst C, Börjeson S, Peterson C. Delayed chemotherapy-induced nausea augmented by high levels of endogeneous noradrenaline . *Brit J Cancer* 1994; 70: 642-45
- 58 Freeman AJ, Cunningham KT, Tyers MB. Selectivity of 5-HT<sub>3</sub>-receptor antagonists and antiemetic mechanisms of action. *Anticancer Drugs* 1992; 3: 79-85
- 59 Fukui HM, Yamamoto T, Ando S, Sasaki S, Sato S. Increase in serotonin levels in the dog ileum and blood by cisplatin as measured by microdialysis . *Neuropharmacology* 1993; 32: 959-68
- 60 Fukui HM, Yamamoto M, Sasaki S, Sato S. Possible involvement of peripheral 5-HT<sub>4</sub> receptors in copper sulfate – induced vomiting in dogs. *Eur J Pharmacol* 1994; 257: 47-52
- 61 Gaedigk A, Blum M, Gaedigk R, Eichelbaum M, Meyer UA. Deletion of the entire cytochrome P450 CYP2D6 gene as a cause of impaired drug metabolism in poor metabolizers of the debrisoquine/sparteine polymorphism. *Am J Hum Genet* 1991; 48: 943-59
- 62 Gleiter CH, Gundert-Remy U. Gender differences in pharmacokinetics. *Eur J Drug Metab Pharmacokinetics* 1996; 21: 123–28
- 63 Goedehals L, Heron JF, Kleisbauer JP, Pagani O, Sessa C. Control of delayed nausea with granisetron plus dexamethasone or dexamethasone alone in patients receiving highly emetogenic chemotherapy: a double blind, placebo-controlled, comparative study. *Ann Oncol* 1998; 9: 661-66
- 64 Gralla RJ. Current issues in the management of nausea and vomiting. *Ann Oncol* 1993; 4 (Suppl.3): S3-S7
- 65 Gralla RJ, Clark RA, Kris MG, Tyson LB. Methodology in anti-emetic trials . *Europ J Cancer* 1991; 27 (Suppl 1): S5-S8
- 66 Gralla RJ, Navari RM, Hesketh PJ. Single-dose oral granisetron has equivalent antiemetic efficacy to intravenous ondansetron for highly emetogenic cisplatin-based chemotherapy. *J Clin Oncol* 1998; 16: 1568-73
- 67 Gralla RJ, Osoba D, Kris MG. Recommendations for the use of antiemetics: evidence-based, clinical tractic guidelines. *J Clin Oncol* 1999; 17: 2971-94

68 Graves T. Emesis as a complication of cancer chemotherapy : pathophysiology, importance and treatment. *Pharmacotherapy* 1992; 12: 337-45

69 Gregory RE, Ettinger DS. 5-HT<sub>3</sub> receptor antagonists for the prevention of chemotherapy-induced nausea and vomiting. A comparison of their pharmacology and clinical efficacy. *Drugs* 1998; 55: 173-89

70 Griese EK, Zanger UM, Brudermans U et al. Assessment of the predictive power of genotypes for the in-vivo catalytic function of CYP2D6 in a German population. *Pharmacogenetics* 1998; 8: 15-26

71 Guzey C, Norstrom A, Spigset O. Change from the CYP2D6 extensive metabolizer to the poor metabolizer phenotype during treatment with bupropion. *Ther Drug Monit* 2002; 24 (3): 436–37

72 Hainsworth JD. Development of serotonin antagonists for the control of chemotherapy – induced emesis. *Semin Surg Oncol* 1993; 9: 279-84

73 Harris RZ, Benet LZ, Schwarz JB. Gender effects in pharmacokinetics and pharmacodynamics. *Drugs* 1995; 50: 222–39

74 Hesketh PJ. Treatment of chemotherapy-induced emesis in the 1990s : improvement by the 5-HT<sub>3</sub> receptor antagonists. *Support Care Cancer* 1994; 2 (5): 286–92

75 Hesketh PJ. Defining the emetogenicity of cancer chemotherapy regimens : relevance to clinical practice. *Oncologist* 1999; 4 (3): 191–96

76 Hesketh PJ. Comparative review of 5-HT<sub>3</sub> receptor antagonists in the treatment of acute chemotherapy-induced nausea and vomiting. *Cancer Invest* 2000; 18 (2): 163-73

77 Hesketh PJ. Potential role of the NK1 receptor antagonists in chemotherapy-induced nausea and vomiting. *Support Care Cancer* 2001; 9: 350-54

78 Hesketh PJ, Gandara DR. Serotonin antagonists: a new class of antiemetic agents. *J Natl Cancer Inst* 1991; 83: 613-20

79 Hesketh PJ, Kris MG, Grunberg SM et al. Proposal for classifying the acute emetogenicity of cancer chemotherapy. *J Clin Oncol* 1997; 15 (1): 103-09

- 80 Hesketh PJ, Van Belle S, Aapro M et al. Differential involvement of neurotransmitters through the time course of cisplatin-induced emesis as revealed by therapy with specific receptor antagonists. *Eur J Cancer* 2003; 39: 1074-80
- 81 Hoyer D, Clarke DE, Fozard JR et al. VII. International Union of pharmacology classification of receptors for 5-Hydroxytryptamine (serotonin). *Pharmacol Rev* 1994; 46: 157-203
- 82 Ingelman-Sundberg M, Oscarsson M, Mc Lellan RA. Polymorphic human cytochrome P450 enzymes: an opportunity for individualized drug treatment. *Trends Pharmacol Sci* 1999; 20: 342-49
- 83 Italian Group for Antiemetic Research: Dexamethasone, granisetron or both for the prevention of nausea and vomiting during chemotherapy for cancer. *N Engl J Med* 1995; 332: 1-5
- 84 Italian Group for Antiemetic Research: double – blind , dose – finding study of four intravenous doses of dexamethasone in the prevention of cisplatin-induced acute emesis. *J Clin Oncol* 1998; 16: 2937-42
- 85 Italian Group for Antiemetic Research : Dexamethasone alone or combined with ondansetron in the prevention of delayed nausea and vomiting induced by chemotherapy. *N Engl J Med* 2000; 342: 1554-59
- 86 Johansson I, Lundquist E, Bertilsson L, Dahl ML, Sjöqvist F, Ingelman-Sundberg M. Inherited amplification of an active gene in the cytochrome P450 CYP2D locus as a cause of ultrarapid metabolism of debrisoquine. *Proc Nat Acad Sci* 1993; 90: 11825-29
- 87 Johansson I, Oscarsson M, Yue QY, Bertilsson F, Sjöqvist F, Ingelman-Sundberg M. Genetic analysis of the Chinese cytochrome P450D locus: characterization of variant CYP2D6 genes present in subjects with diminished capacity for debrisoquine hydroxylation. *Mol Pharmacol* 1994; 46: 452-59 (Abstract)
- 88 Junker A. Antiemetika bei Übelkeit und Erbrechen, induziert durch Zytostatika. *Krankenhauspharmazie* 1999; 5: 170-76
- 89 Junker A, Feyerabend T, Wiedemann GJ, Illiger HJ. Zytostatika- oder Strahlentherapie-induzierte Übelkeit und Erbrechen. *Onkologie* 2000; 6: 244-53
- 90 Kagimoto M, Heim M, Kagimoto K, Zeugin T, Meyer UA. Multiple mutations of the human cytochrome P450IID6 gene (CYP2D6) in poor metabolizers of debrisoquine. *J Biol Chem* 1990; 265: 17209-214



- 91 Kaiser R, Sezer O, Papies A, Bauer S, Schelenz C, Tremblay BB, Possinger K, Roots I, Brockmüller J. Patient-tailored antiemetic treatment with 5-hydroxytryptamine type 3 receptor antagonists according to Cytochrome P450 2D6 genotypes. *J Clin Oncol* 2002; 20: 2805-11
- 92 Kharasch ED, Russell M, Garton K, Lentz G, Bowdle TA, Cox K. Assessment of cytochrome P4503A4 activity during the menstrual cycle using alfentamil as a noninvasive probe. *Anaesthesiology* 1997; 87: 26-35
- 93 Kimura S, Umeno M, Skoda RC, Meyer UA, Gonzalez FJ. The human debrisoquine 4-hydroxylase (CYP2D) locus: sequence and isolation of the polymorphic CYP2D6 gene, a related gene and a pseudogene. *Amer J Human Genet* 1989; 45: 889-904
- 94 Kris MG, Gralla RJ, Tyson LB, Clark RA, Cirrincione C, Groshen S. Controlling delayed vomiting: double-blind, randomised trial comparing placebo , dexamethasone alone and metoclopramide plus dexamethasone in patients receiving cisplatin. *J Clin Oncol* 1989; 7: 108-14
- 95 Kris MG, Radford JE, Pizzo BA, Inabinet R, Hesketh A, Hesketh PJ. Use of NK1 receptor antagonist to prevent delayed emesis after cisplatin. *J Natl Canc Inst* 1997; 89: 817-18
- 96 Kris MG, Roila F, De Mulder PHM, Marty M. Review article : Delayed emesis following anticancer chemotherapy. *Supportive Care Cancer Abstract Volume* 1998; 6: 228-32
- 97 Kutz K. Pharmacology, toxicology and human pharmacokinetics of tropisetron . *Ann Oncol* 1993; 4 (Suppl 3): 15–18
- 98 Laine K, Tybring G, Hartter S, Andersson K, Svensson JO, Widen J, Bertilsson L. Inhibition of cytochrome P4502D6 activity with paroxetine normalizes the ultrarapid metabolizer phenotype as measured by nortriptyline pharmacokinetics and the debrisoquine test. *Clin Pharmacol Ther* 2001; 70: 327–35
- 99 Legrand-Andreoletti M, Stucker I, Marez D et al. Cytochrome P450 (CYP2D6) gene polymorphism and lung cancer susceptibility in Caucasians. *Pharmacogenetics* 1998; 8: 7-14
- 100 Lofters WS, Pater JL, Zee B. Phase III double-blind comparison of dolasetron mesylate and ondansetron and an evaluation of the additive role of dexamethasone in the prevention of acute and delayed nausea and vomiting due to moderately emetogenic chemotherapy. *J Clin Oncol* 1997; 15: 2966-73

101 Lovlie R, Daly AK, Molven A, Idle JR, Steen VM. Ultrarapid metabolizers of debrisoquine: characterization and PCR-based detection of alleles with duplication of the CYP2D6 gene. *FEBS Letters* 1996; 392: 30-34

102 Lundqvist E, Johansson I, Ingelman-Sundberg M. Genetic mechanisms for duplication and multiduplication of the human CYP2D6 gene and methods for detection of duplicated CYP2D6 gene. *Gene* 1999; 226: 327-38

103 Mantovani G, Maccio A, Carreli L. Comparison of oral 5-HT<sub>3</sub>-receptor antagonists and low dose oral metoclopramide plus i.m. dexamethasone for the prevention of delayed emesis in head and neck cancer patients receiving high-dose cisplatin. *Oncol Rep* 1998; 5: 273-80

104 Marez D, Legrand M, Sabbagh N, Lo-Guidice JM, Boone P, Broly F. An additional allelic variant of the CYP2D6 gene causing impaired metabolism of sparteine . *Hum Genet* 1996; 97: 668-70

105 Marez D, Legrand M, Sabbagh N et al. Polymorphism of the cytochrome P450 CYP2D6 gene in a European population : characterization of 48 mutations and 53 alleles, their frequencies and evolution. *Pharmacogenetics* 1997; 7: 193-202

106 Marez D, Sabbagh N, Legrand M, Lo-Guidice JM, Boone P, Broly F. A novel CYP2D6 allele with an abolished splice recognition site associated with the poor metabolizer phenotype. *Pharmacogenetics* 1995; 5: 305-11

107 Martoni S, Angelelli B, Guaraldi M, Strocchi E, Pannuti F. Granisetron (GRA) vs. ondansetron (OND) in the prevention of cisplatin-induced emesis: an open randomized cross-over study (abstract). *Am Soc Clin Oncol* 1993; 13: 431

108 Mc Elroy S, Sachse C, Brockmüller J et al. CYP2D6 genotyping as an alternative to phenotyping for determination of metabolic status in a clinical trial setting. *AAPS Pharm Sci* 2000; 2: E33

109 Mc Lellan RA, Oscarsson M, Seidegard J, Evans DA, Ingelman-Sundberg M. Frequent occurrence of CYP2D6 gene duplication in Saudi Arabians. *Pharmacogenetics* 1997; 7: 187-91

110 Meibohm B, Beierle I, Derendorf H. How important are gender differences in pharmacokinetics ? *Clin Pharmacokinet* 2002; 41: 329-42

111 Milano S, Simon C, Grelot L. In vitro release and tissue release of ileal serotonin after cisplatin induced emesis in the cat. *Clinics in Autonomic Res* 1991; 1: 275- 80

- 112 Miller AD. Central mechanisms of vomiting. *Dig Dis Sci* 1999; 44 (Suppl.): 39-43
- 113 Miller AD, Leslie RA. The area postrema and vomiting. *Front Neuroendocrinology* 1994; 15: 301–20
- 114 Morrow GR, Dobkin PL. Anticipatory nausea and vomiting in cancer patients undergoing chemotherapy treatment: Prevalence , etiology, and behavioural interventions . *Clin Psychol Rev* 1988; 8: 517-36
- 115 Morrow GR, Hickok JT, Rosenthal SN. Progress in reducing nausea and emesis. Comparisons of ondansetron (zofran), granisetron (kytril) and tropisetron (navoban) *Cancer* 1995; 76: 343-57
- 116 Navari RM, Gandara DR , Hesketh PJ. Comparative clinical trial of granisetron and ondansetron in the prophylaxis of cisplatin-induced emesis. *J Clin Oncol* 1995; 13: 1242-48
- 117 Navari RM. Use of placebos in delayed-emesis studies. *J Clin Oncol* 1999; 17: 1644-49
- 118 Navari R, Gralla RJ, Hesketh P. L-754,030, a selective Neurokinin-1 antagonist, reduces cisplatin-induced acute and delayed emesis : a double blind, randomized trial . *Proc Am Soc Clin Oncol* 1998; 17: 197-207
- 119 Navoban Product Information; Novartis Pharma , Basel , Switzerland
- 120 Nitta Y, Nishibori M, Iwagaki H et al. Changes in serotonin dynamics in the gastrointestinal tract of colon-26-tumour-bearing mice: effects of cisplatin treatment. *Naunyn Schmiedeberg`s Arch Pharmacol* 2001; 364: 329-34
- 121 Perez EA. Review of the preclinical pharmacology and comparative efficacy of 5-hydroxytryptamine-3 receptor antagonists for chemotherapy-induced emesis. *J Clin Oncol* 1995; 13: 1036-43
- 122 Perez EA. 5-HT<sub>3</sub> antiemetic therapy for patients with breast cancer. *Breast Cancer Res Treat* 1999; 57 (2): 207-14
- 123 Perez EA, Hesketh PJ, Sandbach EJ. Comparison of single-dose oral granisetron versus intravenous ondansetron in the prevention of nausea and vomiting induced by moderately emetogenic chemotherapy : a multicenter , double-blind , randomized parallel study. *J Clin Oncol* 1998; 16: 754-60

- 124 Possinger K, Späth-Schwalbe E, Kaiser R. Antiemetische Behandlung bei zytostatischer Chemotherapie. *Dtsch Ärzteblatt* 2001; 98: 924–27
- 125 Rademaker M. Do women have more adverse drug reactions ? *Am J Clin Dermatol* 2001; 2: 349–51
- 126 Roh HK, Dahl ML, Johansson J, Ingelman-Sundberg M, Cha YN, Bertilsson L. Debrisoquine and S-Mephenytoin hydroxylation phenotypes and genotypes in a Korean population. *Pharmacogenetics* 1996; 6: 441-47
- 127 Roila F, Del Favero A. Antiemetics revisited. *Curr Opin Oncol* 1997; 9 (4): 321-26
- 128 Roila F, Palladino MA, Ciccarese G, Basurto C. Recent improvements in antiemetic therapy. *Tumori* 1993; 83 (Suppl.2): S3-S14
- 129 Roila F, Tonato M, Basurto I. Antiemetic activity of high doses of metoclopramide combined with methylprednisolone versus metoclopramide alone in cisplatin-treated cancer patients: a randomised double-blind trial of the Italian Oncology Group for Clinical Research. *J Clin Oncol* 1997; 5: 141-49
- 130 Roila F, Tonato M, Cognetti F, Cortesi E, Favalli G, Marangolo M. Prevention of cisplatin-induced emesis: a double-blind multicenter randomized crossover study comparing ondansetron and ondansetron plus dexamethasone. *J Clin Oncol* 1991; 9: 675-78
- 131 Ruff P, Paska W, Goedehals L. Ondansetron compared with granisetron in the prophylaxis of cisplatin-induced acute emesis: a multicentre double-blind, randomized, parallel-group study. *Oncology* 1994; 51: 113–18
- 132 Sachse C, Brockmöller J, Bauer S, Reum T, Roots I. A rare insertion of T 226 in exon 1 of CYP2D6 causes a frameshift and is associated with the poor metabolizer phenotype : CYP2D6\*15. *Pharmacogenetics* 1996; 6 (3): 269-72
- 133 Sachse C, Brockmöller J, Bauer S, Roots I. Cytochrome P4502D6 variants in a Caucasian population: allele frequencies and phenotypic consequences. *Am J Human Genet* 1997; 60 (2): 284–95
- 134 Sachse C, Brockmöller J, Hildebrand M, Müller K, Roots I. Correctness of prediction of the CYP2D6 phenotype confirmed by genotyping 47 intermediate and poor metabolizers of debrisoquine. *Pharmacogenetics* 1998; 8: 181–85

- 135 Sanwald P, David M, Dow J. Characterization of the cytochrome P450 enzymes involved in the in vitro metabolism of dolasetron. Comparison with other indole-containing 5-HT<sub>3</sub> antagonists. *Drug Metab Dispos* 1996; 24: 603-09
- 136 Schwab M, Marx C, Zanger U, Eichelbaum M, Fischer-Bosch M. . Pharmakogenetik der Cytochrom P450 – Enzyme: Bedeutung für Wirkungen und Nebenwirkungen von Medikamenten. *Dtsch Ärzteblatt* 2002; 99: 497-504
- 137 Schwörer K, Racke K, Kilbinger H. Cisplatin increases the release of 5-hydroxytryptamine (5-HT) from the isolated vascularity perfused small intestine of the guinea-pig : involvement of 5-HT<sub>3</sub> receptors. *Naunyn Schmiedeberg's Arch Pharmacol* 1991; 344: 143-49
- 138 Seynaeve C, De Mulder PHM, Verweij J. Pathophysiology of cytotoxic drug-induced emesis : Far from crystal-clear. *Pharm Weekl Sci* 1990; 13: 1-6
- 139 Smyth J F. Delayed emesis after high-dose cisplatin – the residual problem. *Proc Eur Soc Med Oncol* 1992; Nov 8, Lyon France
- 140 Sorbe B. Tropisetron in the prevention of chemotherapy-induced nausea and vomiting: the Nordic experience. *Ann Oncol* 1993; 4: 39-42
- 141 Steijns LSW, Van der Weide J. Ultrarapid drug metabolism: PCR based detection of CYP2D6 gene duplication. *Clin Chem* 1998; 44: 914–17
- 142 Steward DJ. Cancer therapy, vomiting and antiemetics. *Can J Physiol Pharmacol* 1990; 68: 304-13
- 143 Stewart A, Mc Quade B, Cronje JDE. Ondansetron compared with granisetron in the prophylaxis of cyclophosphamide-induced emesis in out-patients: a multicenter, double-blind, double-dummy randomized, parallel-group study. *Oncology* 1995; 52: 202–10
- 144 Tanaka E. Gender-related differences in pharmacokinetics and their clinical significance. *J Clin Pharm Ther* 1999; 24: 339–46
- 145 Tavorath R, Hesketh PJ. Drug treatment of chemotherapy-induced delayed emesis. *Drugs* 1996; 52 (5): 639–48
- 146 Tonato M, Roila F, Del Favero A. Methodology of antiemetic trials: a review . *Ann Oncol* 1991; 2: 107–14
- 147 Tonato M, Roila F, Del Favero A, Ballatori E. Antiemetics in cancer chemotherapy – historical perspective and current state of the art. *Cancer* 1994; 2: 150–60

- 148 Tonato M, Roila F, Del Favero A. Are there differences among the serotonin antagonists ? Support Care Cancer 1994; 2: 293-96
- 149 Tortorice PV, O'Connell MB. Management of chemotherapy-induced nausea and vomiting. Pharmacotherapy 1990; 10: 129-45
- 150 Tremblay PB, Kaiser R, Sezer O, Rösler N, Schelenz C, Possinger K, Roots I, Brockmüller J. Variations in the 5-hydroxytryptamine type 3 B receptor gene as predictors of the efficacy of antiemetic treatment in cancer patients. J Clin Oncol 2003; 21: 2147–55
- 151 Tyers MB. Pharmacology and preclinical antiemetic properties of ondansetron Semi Oncol 1992; 4 (Suppl 10): 1–8
- 152 Tyers MB, Freeman AJ. Mechanisms of antiemetic activity of 5-HT<sub>3</sub> antagonists. Oncology 1991; 49: 263-68
- 153 Villikka K, Kivisto KT, Neuvonen PJ. The effect of rifampin on the pharmacokinetics of oral and intravenous ondansetron. Clin Pharmacol Ther 1999; 65 (4): 377-81
- 154 Waldvogel HH. Antiemetische Therapie. Thieme-Verlag Stuttgart 1995
- 155 Watson JW, Gonsalves SF, Fossa AA, Mc Lean S, Seeger T, Obdach S, Andrews PLR. The antiemetic effects of CP-99,994 in the ferret and the dog: role of the NK<sub>1</sub> receptor. Brit J Pharmacol 1995; 115: 84-94
- 156 Wickham R. Nausea and vomiting. In : Yarbo CH, Frogge MH, Goodman M, eds: Cancer Symptom Management. 2<sup>nd</sup> Edition, Sudbury, Mass.: Jones and Bartlett Publishers 1999, 228-63
- 157 Wilder-Smith OHG, Borgeat A, Chappuis P, Fathi M, Forni M. Urinary serotonin metabolite excretion during cisplatin chemotherapy. Cancer 1993; 72: 2239-41
- 158 Wolf H. Preclinical and clinical pharmacology of the 5-HT<sub>3</sub> receptor antagonists. Scand J Rheumatol Suppl. 2000; 113: 37-45
- 159 Yalcin S, Tekuzman G, Baltali E. Serotonin receptor antagonists in prophylaxis of acute and delayed emesis induced by moderately emetogenic, single-day chemotherapy : a randomized study. Amer J Clin Oncol 1999; 22: 94-96

160 Yamakuni H, Sawai H, Maeda Y et al. Probable Involvement of the 5-HT<sub>4</sub>-Receptor in Methotrexat-Induced Delayed Emesis in Dogs. *Pharmacology* 2000; 292: 1002-07

161 Zofran (ondansetron hydrochloride) Injection and Tablets Product Information . Glaxo Wellcome Oncology/HIV. In: *Physician`s Desk Reference* . 52<sup>nd</sup> ed. , Montvale,NJ Medical Economics Company 1998, 1177-83