

# Ausgewählte pharmakogenetische Studien in verschiedenen Anwendungsbereichen

### Pharmakogenetik in diversen Indikationen

- → Swen Jesse J et al., The Lancet 2023 Feb 401;347-56,
   A 12-gene pharmacogenetic panel to prevent adverse drug reactions: an open-label, multicentre, controlled, clusterrandomised crossover implementation study (6.944 patients)
  - Pharmacogenetics-guided prescribing results in a **30% reduction** of clinically relevant adverse drug reactions."

## Pharmakogenetik in der Psychiatrie

- → Skokou Maria et al., The Lancet 2024; 101: 105009
   Clinical implementation of preemptive pharmacogenomics in psychiatry (1.076 patients)
  - "PGx-guided arm present with 34.1% less adverse drug reactions compared to control arm, 41.2% less hospitalisations and 40.5% less re-admissions, less duration of initial hospitalisations and duration of hospitalisation upon readmission."
- → Vos F. Cornelis et al, JAMA Network Open, 2023; 6(5): 12443
   Effectiveness of Genotype-Specific Tricyclic Antidepressant Dosing in Patients With Major Depressive Disorder A Randomized Clinical Trial (111 patients)
  - ➢ "In this randomized clinical trial, pharmacogenetics-informed treatment resulted in faster attainment of therapeutic TCA concentrations, with potentially fewer and less severe adverse effects."
- → Bousman CA, Arandjelovic K, Mancuso SG, Eyre HA, Dunlop BW., Pharmacogenomics. 2019;20(1):37–47 Pharmacogenetic tests and depressive symptom remission: a meta-analysis of randomized controlled trials (1.737 patients)
  - Pharmacogenetic-guided therapy **1.71 times more likely to achieve symptoms remission** relative to individuals who received usual treatment."
- → Altar CA et al., Mol Neuropsychiatry 2015;1:145–155 Clinical Utility of Combinatorial Pharmacogenomics-Guided Antidepressant Therapy: Evidence from Three Clinical Studies (258 patients)

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#### Pharmakogenetik in der Kardiologie

- → Claassens DMF et al., New Engl J Med 2019, 381;17
   A Genotype-Guided Strategy for Oral P2Y12 Inhibitors in Primary PCI (2.488 patients)
  - "CYP2C19 genotype-guided strategy for selection of Clopidogrel therapy was noninferior to standard treatment with ticagrelor or prasugrel at 12 months with respect to thrombotic events and resulted in a **lower incidence of bleeding**."

## Pharmakogenetik in der Onkologie

- → He W et al, Ann. Oncol. 2021 Oct;32(10):1286-1293 CYP2D6 genotype predicts tamoxifen discontinuation and drug response: a secondary analysis of the KARISMA trial (1440 patients)
  - "Poor CYP2D6 metabolizers are likely to experience an impaired response to tamoxifen, measured through mammographic density reduction. In contrast, ultrarapid CYP2D6 metabolizers are at risk for exaggerated response with pronounced adverse effects that may lead to treatment discontinuation."
- → Deenen MJ, et al., J Clin Oncol. 2016; 34(3):227–34.
  Upfront Genotyping of DPYD\*2A to Individualize Fluoropyrimidine Therapy: A Safety and Cost Analysis (2.038 patients)
  - The risk of 5-FU-induced toxicity was significantly **reduced from 73%** in historical controls (n =48) **to 28% by genotype-guided dosing** (P < .001); drug-induced death was reduced from 10% to 0%."

#### Pharmakogenetik in der Schmerzbehandlung

- → Smith DM, et al., Genet Med. 2019;21(8):1842 CYP2D6-guided opioid therapy improves pain control in CYP2D6 intermediate and poor metabolizers: a pragmatic clinical trial (375 patients)
  - "Within the same subgroup of IM/PMs prescribed tramadol or codeine at baseline, CYP2D6-guided group experienced a 30% reduction in composite pain intensity compared with the usual care group."

## Pharmakogenetik in der Polypharmazie

- → Hayashi M et al., es Social Adm Pharm 2021 Aug 20;S1551-7411(21)00313-2, Applications for pharmacogenomics in pharmacy practice: A scoping review (43 PGx studies reviewed)
  - "Pharmacogenetic multi-gene panel tests within MTM (medication therapy management) demonstrate an approximate 50% decrease in emergency visits and hospitalizations in elderly polypharmacy patients."

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