

# Dextromethorfan + Panobinostat

# M1323

| Onderbouwend   | Stof                             | Effect  | Code |
|--|----------------------------------|---|------|
| Feld R.<br>Cancer Chemother Pharmacol 2013;72:747-55.<br>doi: 10.1007/s00280-013-2237-3.   | dextromethorfan + panobinostat   | dextromethorfan ↑AUC <sub>0-∞</sub> 1.64x en Cmax 1.83x; t1/2 ongewijzigd<br>dextromethorfan metabolism ↑AUC <sub>0-∞</sub> 1.3x, Cmax vrijwel ongewijzigd (1.05x) ; t1/2 ongewijzigd<br>Regime: dextromethorfan 60 mg op dag 1, panobinostat 20 mg op dag 3+5, en combinatie op dag 8; 14 patients with advanced cancer who have functional CYP2D6 genes.<br>Auteurs: panobinostat weakly inhibited a sensitive CYP2D6 substrate in cancer patients by increasing DM exposure by less than twofold. Safety monitoring of sensitive CYP2D6 substrates with narrow therapeutic index is recommended when co-administering with panobinostat in future clinical practice. | 3A   |
| SPC + EPAR Farydak   | dextromethorfan + panobinostat   | getallen uit Feld 2013.<br>dextromethorfan ↑AUC 1.6x en Cmax 1.8x<br>Study B2109: In the core phase of the study, panobinostat 20 mg was administered once per day on days 3, 5, and 8. Oral dextromethorphan at 60 mg was administered on the mornings of Day 1 and Day 8. There was a high inter-subject variability in dextromethorphan pharmacokinetics even though only extensive CYP2D6 subjects were selected.   | 2A   |
| Farydak Prescribing Information<br><a href="http://www.accessdata.fda.gov/drugsatfda_docs/label/2015/205353s000lbl.pdf">http://www.accessdata.fda.gov/drugsatfda_docs/label/2015/205353s000lbl.pdf</a> .<br>Geraadpleegd 12-11-2015. | CYP2D6-substraten + panobinostat | zelfde info.<br>Avoid combination with sensitive CYP2D6 substrates (i.e., atomoxetine, dextromethorphan, metoprolol, nebivolol, perphenazine, tolterodine, venlafaxine) or CYP2D6 substrates that have a narrow therapeutic index (i.e., thioridazine, pimozide). If concomitant use of CYP2D6 substrates is unavoidable, monitor patients frequently for adverse reactions.  | 2A   |

| Overig      | Stof                             | Effect  |
|-------------|----------------------------------|---|
| SPC Farydak | CYP2D6-substraten + panobinostat | Vermijd panobinostat bij gebruik van CYP2D6-substraten met kleine therapeutische index (inclusief maar niet beperkt tot pimozide).<br>Bij combinatie met gevoelige CYP2D6-substraten moeten de doses van afzonderlijke CYP2D6-substraten worden getitreerd op basis van verdraagbaarheid en moeten patiënten regelmatig gecontroleerd worden op bijwerkingen. |

## Opmerkingen

PubMed: niets op andere CP2D6-substraten.

Stockley: -

|             |  |
|-------------|--|
| Risicogroep |  |
|-------------|--|

| Interactie     | Actie | Datum |
|----------------|-------|-------|
| Beslissing WFG | Ja    | Nee   |