

Using the NZF Interactions Checker

The New Zealand Formulary (NZF) contains a function that identifies potential drug interactions and their clinical significance. The Interactions Checker allows a user to search for potential interactions between any number of medicines, and between medicines and some other substances that are known to significantly interact, e.g. ethanol, tobacco, grapefruit and some complementary and Chinese herbal medicines.

The NZF Interactions Checker is located at the top of each page of the NZF online, beneath the blue NZF banner and beside the “Search NZF” box at the top of the screen. The Interactions and the Search NZF boxes are either grey or white depending on which is “active” (white) at the time (Figure 1).

To use the Interactions Checker, click on the “Interactions” box. A brief explanation of the Interactions Checker will then appear. A search box at the top of the page is labelled “Enter medicines”. Type the first few letters of a generic or brand name and select the medicine you want from the drop down box that appears. Search for potential interactions by adding medicines to the search box one by one. If only one medicine is chosen, a list of all the known interactions of that medicine will appear. There is no maximum number of medicines that can be entered into the Interactions Checker each time a query is performed.

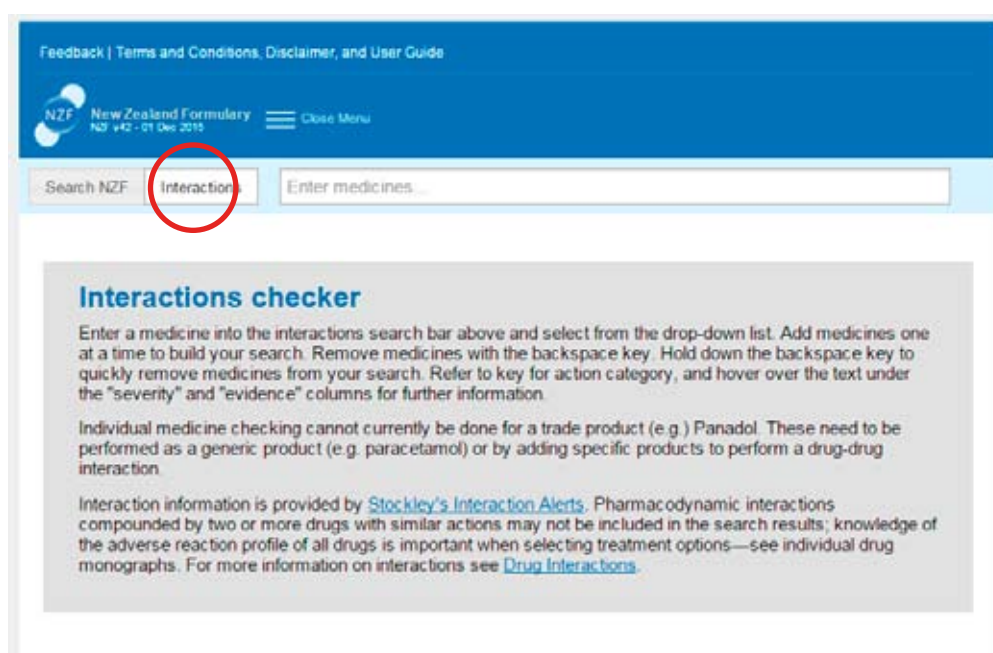


Figure 1: Interactions search box shown in NZF online.

The results of the interactions search are displayed in a table format, with the most clinically significant interactions appearing at the top of the table (Figures 2 and 3).

The results table displays:

1. The sets of medicines that interact, from the list that has been entered
2. A brief explanation of the mechanism for an interaction, if known
3. The interactions identified in order of decreasing clinical significance, with a colour-coded square to indicate what action is required: red = Avoid, orange = Adjust, gold = Monitor, green = Information (for less significant interactions that require monitoring depending on patient characteristics), white = No action
4. Specific advice on what action to take, e.g. avoid combination, alter dose(s), monitor for adverse effects, monitor for reduced effect, monitor plasma concentration
5. Expected severity of the interaction if adjustments to treatment are not made
6. The level of evidence for the combination being deemed to be a drug interaction and for identifying the interaction's clinical importance: Extensive evidence, Formal study, Case reports, Theoretical evidence

Search NZP Interactions

Please add additional medicines to the interactions search bar above to perform a drug-drug interaction check.

Interactions between: cilazapril; hydrochlorothiazide; furosemide; metoprolol succinate KEY: ■ avoid ■ adjust ■ monitor ■ information ■ no action

Medicines	Explanation	Action	Severity	Evidence
cilazapril (systemic) and furosemide (systemic)	The concurrent use of an ACE inhibitor with a loop diuretic is normally safe and clinically beneficial; however, first dose hypotension can occur, particularly if the dose of diuretic is high. Rare cases describe renal impairment. Diuretic-induced hypokalaemia can still occur on concurrent use.	■ Start the ACE inhibitor at the lowest dose. Advise patients to lie down if dizziness, lightheadedness, etc. occurs. Furosemide 80 mg daily or more (or equivalent); monitor closely, consider stopping the diuretic 24 hours before starting the ACE inhibitor, or monitor for 2 hours or until the blood pressure is stable.	Moderate	Formal study
cilazapril (systemic) and hydrochlorothiazide (systemic)	The concurrent use of an ACE inhibitor with a thiazide is normally safe and clinically beneficial; however, first dose hypotension can occur, particularly if the dose of diuretic is high. Rare cases describe renal impairment. Diuretic-induced hypokalaemia can still occur on concurrent use.	■ Start the ACE inhibitor at the lowest dose. Advise patients to lie down if dizziness, lightheadedness, etc. occurs. Furosemide 80 mg daily or more (or equivalent); monitor closely, consider stopping the diuretic 24 hours before starting the ACE inhibitor, or monitor for 2 hours or until the blood pressure is stable.	Moderate	Formal study
cilazapril (systemic) and metoprolol (systemic)	Concurrent use is generally well tolerated and can be clinically beneficial. The blood pressure reduction is enhanced, as would be expected.	■ No action needed, unless blood pressure-lowering effects become excessive.	Nothing expected	Theoretical

Figure 2: Example of interactions table of results for four selected medicines

Search NZP Interactions

Please add additional medicines to the interactions search bar above to perform a drug-drug interaction check.

Interactions between: warfarin; st john's wort; ethanol KEY: ■ avoid ■ adjust ■ monitor ■ information ■ no action

Medicines	Explanation	Action	Severity	Evidence
warfarin (systemic) and st john's wort (systemic)	St John's wort (Hypericum perforatum) can reduce the anticoagulant effects of warfarin.	■ Monitor the INR on concurrent use and adjust the coumarin dose if necessary. Note that the CSM in the UK advise stopping St John's wort.	Moderate	Case reports
warfarin (systemic) and alcohol (systemic)	Heavy drinkers of alcohol may have a reduced response to coumarins. Binge drinking may increase the risk of bleeding. Moderate alcohol intake is unlikely to alter the response to coumarins.	■ Heavy drinkers of alcohol may need above-average doses of a coumarin; however, avoid anticoagulation unless heavy or binge drinkers stop drinking alcohol. Counsel patients about moderate alcohol intake when they start an anticoagulant.	Nothing expected	Formal study

Figure 3: Example of interactions table of results for one medicine and two interacting substances. N.B. the search term "ethanol" must be used, rather than alcohol, when entering substances in the interactions checker. "Tobacco" is the correct search term for identifying interactions between medicines and smoking.

The drug interactions information in the NZF Interactions Checker is provided by “Stockley’s Interaction Alerts” which is a United Kingdom source of interactions information that is easily applied and relevant to clinical decision making in New Zealand. Stockley’s Interaction Alerts holds interaction information on all medicines that appear in the NZF.

The NZF Interactions Checker does not include opposing or additive pharmacodynamic effects in the definition of an interaction. Clinicians therefore need to be alert for opposing or additive pharmacological or adverse effects between medicines that may not be identified as an interaction by the NZF Interactions Checker.

Examples of such combinations of medicines that are not identified as significant drug interactions in the NZF Interactions Checker include:

- An oral non-selective beta-blocker such as carvedilol, labetalol or propranolol and an inhaled beta-1 receptor agonist such as salbutamol, which should not be taken together as their effects oppose each other
- Omeprazole, a proton-pump inhibitor, and ranitidine, a histamine-2 receptor antagonist, are both indicated for the treatment of gastric ulceration but do not usually provide additional benefit if used together

Additional interactions information appears in each individual NZF drug monograph under the heading “Interactions”. There are links to both the drug interaction summaries from the British National Formulary and to the Stockley’s Interaction Alerts information that appears in the NZF interactions checker (Figure 4). Occasionally supplementary statements are also included in this section.



Figure 4: Example of a drug monograph, showing the location of links to interactions information

To try out the NZF Interactions Checker, visit:

www.nzf.org.nz