

MMV develops framework to assess risk of resistance for antimalarial compounds

22 August 2012

Medicines for Malaria Venture has developed a framework to evaluate the risk of resistance for the antimalarial compounds in its portfolio. A paper based on this work: A framework for assessing the risk of resistance for antimalarials in development has been published in the *Malaria Journal* today.

Resistance defines the longevity of every antiinfective drug, so it is important when developing <u>new medicines</u> for malaria, to check how easily promising antimalarial compounds will select for resistance. Once this is known, it facilitates the prioritization of not only the most efficacious compounds but also the most robust ones.

"By profiling our portfolio as early as possible in terms of resistance liabilities, be they pre-existing or acquired, we are attempting to ensure that none of the compounds will fall to potential resistance," said Tim Wells, Chief Scientific Officer, MMV, and one of the authors of the paper. "This will also help us cost-effectively accelerate the drug development process, and be prepared in advance with a full resistance profile which is required by regulatory authorities before a new drug can be approved."

A cross-<u>resistance test</u> using a panel of multi-drug <u>resistant strains</u> of the parasite will check for preexisting resistance liability. This will ensure that none of MMV's compounds are cross-resistant with other drugs, e.g., if a parasite is resistant to <u>chloroquine</u>, its resistance will also be profiled against MMV compounds.

The framework also includes selection experiments in the laboratory that measure how easy it is for the parasite to develop resistance, in other words, the likelihood of the occurrence of mutations that confer resistance. This is achieved by measuring the minimal inoculum for resistance (MIR) - the minimum number of parasites from which a resistant one is likely to be selected by drug pressure. Although this is already being done, the

framework offers a standard, systematic method.

This new framework will result in a fully profiled portfolio for MMV in terms of resistance and could also be used by other malaria researchers to test their compounds for potential resistance, measure the genetic ability of <u>parasites</u> to develop resistance and the intensity of the resistance.

Provided by Medicines for Malaria Venture



APA citation: MMV develops framework to assess risk of resistance for antimalarial compounds (2012, August 22) retrieved 7 September 2022 from <u>https://medicalxpress.com/news/2012-08-mmv-framework-resistance-antimalarial-compounds.html</u>

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