

# Catechin Isolated from *Garcinia celebica* Leaves Inhibit *Plasmodium falciparum* Growth through the Induction of Oxidative Stress

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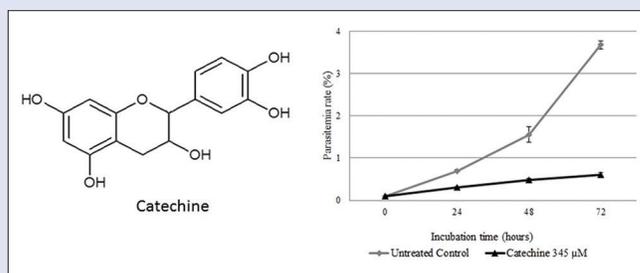
## ABSTRACT

**Background:** Resistance of antimalarial drugs to *Plasmodium falciparum* has become a major concern in malaria eradication. Although it is also affected by several socioeconomic factors, a new antiplasmodial agent is needed for a global malaria control program. **Objective:** In this study, we attempted to uncover the antiplasmodial properties of *Garcinia celebica*, an Indonesian medicinal plant, along with the responsible compound and its possible mechanism. **Materials and Methods:** The *G. celebica* leaves were ethanol extracted and fractionated based on their polarity using *n*-hexane, ethyl acetate, and water. The antiplasmodial activity was tested *in vitro* against chloroquine-resistant *P. falciparum* at 100 µg/ml for 72 h. The active compound of the most active ethyl acetate fraction was subsequently isolated using column chromatography and identified by nuclear magnetic resonance. **Results:** The IC<sub>50</sub> of (+)-catechin, the characterized compound, against *P. falciparum* was 198 µM in 24 h and experiment. The isolated catechin inhibited *P. falciparum* growth in both trophozoite and schizont stages. An additional experiment also suggests that the antiplasmodial property of catechin occurs through the induction of the oxidative stress to *P. falciparum*. **Conclusion:** This result shows that the potential of catechin and its antimalarial properties should be explored further. **Key words:** Catechin, *Garcinia celebica*, malaria, *Plasmodium falciparum*

## SUMMARY

- *Garcinia celebica* leaf extract and fractions inhibit *Plasmodium falciparum* growth
- Catechin, the active compound of *Garcinia celebica* leaf extract, inhibits *Plasmodium falciparum* growth in a time- and dose-dependent manner

- Catechin inhibits *Plasmodium falciparum* growth in both trophozoite and schizont stages, through induction of oxidative stress.



**Abbreviations used:** RBC: Red Blood Cells; IC50: Inhibition Concentration 50; MeOH: Methanol; RPMI: Roswell Park Memorial Institute; EI: Electron Ionization.

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## INTRODUCTION

Malaria is a deadly parasitic disease<sup>[1]</sup> and causes 2 million deaths annually worldwide.<sup>[2]</sup> The biggest problem in its eradication is the growing resistance of *Plasmodium falciparum*, which may cause the most fatal form of malaria,<sup>[3]</sup> to almost all of the currently available malarial drugs.<sup>[4]</sup> Even with the current multidrug therapy program, *P. falciparum* has developed drug resistance against the latest antiplasmodial drug from the artemisinin family.<sup>[5]</sup> Although malaria drug resistance is also affected by several socioeconomic factors, a new antiplasmodial agent is needed for a global malaria control program.

From the hundreds of compounds tested for their antiplasmodial activities, plants and other natural compound are among the most promising.<sup>[6-8]</sup> Quinine, the first antimalarial agent, and the next generation of antimalarial agents, including lapachol and artemisinin, have been isolated from plants.<sup>[9,10]</sup> We previously reported the antiplasmodial activity of kaempferol-3-*O*-rhamnoside isolated from *Schima wallichii* leaves against chloroquine-resistant *P. falciparum*.<sup>[11]</sup> Furthermore, our group also reported the antiplasmodial potential of several selenium compounds to induce apoptosis-like cell death in *P. falciparum*.<sup>[12]</sup>

One of the approaches to find a novel drug from plants is using plants that are consumed by particular groups.<sup>[13]</sup> In recent years, we focused our research on discovering bioactive compounds from plants commonly consumed by nonhuman primates, as they are considered to be a promising source of products applicable for the management of human disease.<sup>[14]</sup> In the present study, we focused on the exploration of the antiplasmodial activities of *Garcinia celebica* leaves, one of the plants that is consumed by nonhuman primates with anticancer potential.<sup>[15]</sup> In our previous studies, we reported the anticancer potential of kaempferol-3-*O*-rhamnoside,<sup>[16,17]</sup> a compound that is

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