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ACCEPTED MANUSCRIPT

Tyrosinase inhibition by some flavonoids: Inhibitory activity, mechanism by in vitro and

in silico studies

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**Abstract** 

Flavonoids are main polyphenolic groups widely distributed to fruits, vegetables and

beverages we consumed daily. They exhibit many biological effects. We tested tyrosinase

inhibitor potential of structurally related (1-9) flavonoids and found that all the tested

materials possessed tyrosinase inhibitory effect compared to the positive control, kojic acid. 2

exhibited the strongest tyrosinase inhibitory effect with an IC<sub>50</sub> value of  $40.94 \pm 0.78 \mu M$  in a

competitive manner. According to kinetic analysis 1, 4 and 7 were found to be competitive

inhibitors, 3, 5, and 6 noncompetitive inhibitors of tyrosinase. According to the docking

studies, A and C ring of the flavonoid structure, hydroxyl substituent at the 7<sup>th</sup> position, and

hydroxyl substituents at para or para and meta position of ring B play key role for

competitive inhibition of the enzyme.

**Keywords:** flavonoid; molecular docking; tyrosinase

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