Effect of curcumin supplementation on exercise-induced muscle damage: a narrative review

K. Nanavati1 · K. Rutherfurd-Markwick2 · S. J. Lee3 · N. C. Bishop4 · A. Ali1

Received: 29 October 2021 / Accepted: 15 June 2022 / Published online: 13 July 2022 © The Author(s) 2022

Abstract
Curcumin, a natural polyphenol extracted from turmeric, is a potent antioxidant and anti-inflammatory agent. In the past few decades, curcumin’s ability to impact chronic inflammatory conditions such as metabolic syndrome, arthritis, and cancer has been widely researched, along with growing interest in understanding its role in exercise-induced muscle damage (EIMD). EIMD impacts individuals differently depending on the type (resistance exercise, high-intensity interval training, and running), intensity, and duration of the exercise. Exercise disrupts the muscles’ ultrastructure, raises inflammatory cytokine levels, and can cause swelling in the affected limb, a reduction in range of motion (ROM), and a reduction in muscular force-producing capacity. This review focuses on the metabolism, pharmacokinetics of various brands of curcumin supplements, and the effect of curcumin supplementation on EIMD regarding muscle soreness, activity of creatine kinase (CK), and production of inflammatory markers. Curcumin supplementation in the dose range of 90–5000 mg/day can decrease the subjective perception of muscle pain intensity, increase antioxidant capacity, and reduce CK activity, which reduces muscle damage when consumed close to exercise. Consumption of curcumin also improves muscle performance and has an anti-inflammatory effect, downregulating the production of pro-inflammatory cytokines, including TNF-α, IL-6, and IL-8. Curcumin may also improve oxidative capacity without hampering training adaptations in untrained and recreationally active individuals. The optimal curcumin dose to ameliorate EIMD is challenging to assess as its effect depends on the curcumin concentration in the supplement and its bioavailability.

Keywords Curcumin · Pharmacokinetics · Inflammation · Muscle soreness · Oxidative stress · Antioxidant

Introduction
Curcumin, chemically known as 1,7-bis(4-hydroxy-3-methoxyphenyl)-1,6-heptadiene-3,5-dione or diferuloylmethane [1], is isolated from the plant Curcuma longa [2]. Over the past few decades, curcumin has been widely researched for its antioxidant and anti-inflammatory properties in numerous chronic and malignant diseases such as metabolic syndrome [3], arthritis [4, 5], and cancer [6–8]. However, there has been a growing interest in understanding the effect of curcumin on exercise-induced muscle damage (EIMD) in recent years. EIMD affects all individuals depending upon the type, intensity, and duration of the exercise they undertake and training status of the individual [9, 10]. Resistance training [11], high-intensity interval training [12], trail running [13, 14], and downhill running [15] contribute to EIMD, leading to ultrastructural muscular disruption and an increase in inflammatory cytokine levels. Swelling of the affected limb, decreased range of motion (ROM),
and impaired muscle force-producing capacity which can result from EIMD are undesirable [16–18].

Curcumin has been shown to attenuate muscle soreness, improve performance, reduce blood levels of inflammatory markers, and enhance endogenous oxidative capacity post-exercise [19–26]. Muscle damage is prominent post-exercise due to the release of prostaglandins under the influence of cyclooxygenase (COX-1 and COX-2), which contributes to redness, swelling, and pain at the site of damage [27, 28]. Curcumin downregulates the expression of COX-2 and thus decreases the release of prostaglandins [29] which in turn reduces muscle damage [19–23]. Exercise, and production of prostaglandins under the influence of COX-2, leads to increased membrane permeability [30, 31] and releases creatine kinase (CK) into the interstitial fluid and then circulation via the lymphatic system [32] and indicates muscle damage. Although research on the chronic effects of curcumin supplementation is limited, data so far have not demonstrated any ergolytic effects as observed following supplementation with other natural antioxidants such as vitamin C and E supplementation. A detailed review [36] on the effects of vitamin C and vitamin E supplementation on exercise performance suggests that these antioxidant supplementations may impair neuromuscular adaptation by affecting muscle mitochondrial biogenesis and muscle hypertrophy [33].

Clinical trials indicate consumption of curcumin close to exercise downregulates cyclooxygenase that influences membrane permeability [31] and offers a membrane-protective effect by altering the structure of the cell membrane to improve its integrity [34]. Curcumin also helps in reducing inflammation by hindering the activation of nuclear factor-kappa B (NF-κB), suppressing the activation and phosphorylation of Janus kinase/signal transducers and activators of transcription (JAK/STAT) proteins, and inhibiting mitogen-activated protein kinase (MAPK) signalling that releases inflammatory markers such as tumour necrosis factor-alpha (TNF-α), interleukin-8 (IL-8), and interleukin-6 (IL-6) at the site of damage [35]. In addition, the downregulation of NF-κB can also lead to the elevation of antioxidant responses by activation of nuclear factor erythroid 2-related factor 2 (NRF2) [36]. NRF2 regulates the synthesis of antioxidant proteins that protect against oxidative damage triggered by injury and inflammation [37].

Curcumin can be tolerated without any associated toxicity at 8000 mg/day [38]. However, poor aqueous solubility [39] and low bioavailability [40] of curcumin have led to the development of different curcumine formulations such as nanoparticles [41], phytosomes [42], micelles [43], and phospholipid complexes [44]. Each formulation contains varying levels of curminoids and has a different rate of absorption, making it difficult to conclude a single recommended dose. Nevertheless, different curcumin formulations have been shown to effectively reduce EIMD and inflammation in doses varying from 90 mg/day to 5000 mg/day [21, 25]. This narrative review evaluates the different curcumin formulations and their effect on EIMD, inflammation, and oxidative markers.

**Methods**

The databases SCOPUS, Medline (PubMed), and Web of Science (WOS) were searched using a mix of Medical Subject Headings (MeSH) and free words for key concepts related to curcumin, muscle, exercise, inflammation, recovery, along with bioavailability of curcumin as follows: (“curcumin” OR “turmeric”) AND (“muscle damage” OR “delay onset muscle soreness” OR “DOMS” OR “inflammation” OR “inflammatory” OR “inflammatory markers” OR “oxidative stress”) AND (“exercise”). For articles on other natural antioxidants and their effect on exercise performance, search terms included (“antioxidants”), (“vitamin E and C”), (“tart cherry juice”), (“natural extracts”) AND (“exercise” OR “exercise-induced muscle damage” OR “exercise performance”), between December 2020 and March 2021. Only full-text articles (written in English) describing human trials were included for review.

**Curcumin metabolism**

The bioavailability of curcumin is low due to its water insolubility and poor metabolism in the small intestine and liver, where it undergoes extensive reductive and conjugative metabolism and is finally eliminated through the gall bladder [45]. The metabolism of curcumin can be divided into two phases (Fig. 1). Phase I comprises the reduction of its double bonds to dihydrocurcumin, tetrahydrocurcumin, hexahydrocurcumin, and octahydrocurcumin by reductases in enterocytes and hepatocytes [40]. For phase II, both curcumin and its metabolites from phase I undergo conjugation with sulphate at its phenolic site in the hepatic and intestinal cytosol. In addition, metabolites also undergo glucuronidation via UDP-glucuronosyltransferase in the intestinal and hepatic microsomes [40]. Alternatively, curcumin can be metabolised by intestinal microbiota, such as *Escherichia coli* to tetrahydrocurcumin with the help of an NADPH-dependent reductase and by *Blautia* sp. MRG-PMF1 (anaerobic bacterial strain) that facilitates curcumin demethylation to form demethylcurcumin and bis-demethylcurcumin [46].

**Pharmacokinetics of curcumin supplements**

Absorption, distribution, hepatic and intestinal metabolism, and excretion regulate the bioavailability of ingested curcumin [45]. In addition, the physicochemical properties of each curcumin formulation and the dose, along with its rate
of degradation in the lumen, lipophilicity, and gastric emptying time, help determine the body’s pharmacokinetics [47]. Some studies have assessed plasma curcumin levels after hydrolysis of blood plasma samples, however, such hydrolysis prior to extraction masks the amount of free, bioactive curcumin and total curcuminoids as compared to non-hydrolysed samples [50]. When treated with the enzymes β-glucuronidase and sulfatase, curcumin generates glucuronide curcumin and curcumin sulphate, which are the primary circulating forms of curcumin but are physiologically inactive conjugates. This obscures the free bioactive curcumin and overestimates the amount of curcumin detected, thus providing incorrect and misleading results regarding the bioavailability of the formulation [48]. This highlights the importance of reporting free curcumin in the plasma without hydrolysis of the sample. Plasma samples obtained from studies involving Theracurmin [41], Meriva [42], NovaSol [43], CurQfen [49], Longvida [44], and Curcumin C3 Complex [50], along with their pharmacokinetic parameters are presented in Tables 1 and 2 [47].

The evidence from human trials suggests that formulating curcumin improves systemic exposure and increases area under curve (AUC) and maximum blood concentration of curcumin, and thus, increases the bioavailability of curcumin ($C_{\text{max}}$) [41–44, 49, 50].

The use of gum ghatti in a water-soluble formulation called Theracurmin [41] led to a preparation of a stable water-soluble complex that contributed to colloidal dispersion and enhanced gastrointestinal absorption [41]. A dose of 30 mg of Theracurmin containing 3.6 mg curcuminoids showed a 27-fold improvement in its AUC (0–6 h), with a $T_{\text{max}}$ of 1 h. Another water-soluble curcumin: NovaSol [43] demonstrated the highest $C_{\text{max}}$ along with 185-fold better bioavailability compared to its native form, with a single dose of 500-mg of curcuminoids. The improved relative bioavailability was contributed by the micellar-based curcumin formulation with Tween 80 (non-ionic surfactant and

---

**Fig. 1** Metabolic pathways of curcumin

---

**Overview of bioavailability of different curcumin formulations**

Information on dosage, study type, and study population of six different formulations, namely Theracurmin [41], Meriva [42], NovaSol [43], CurQfen [49], Longvida [44], and Curcumin C3 Complex [50], along with their pharmacokinetic parameters are presented in Tables 1 and 2 [47]. The evidence from human trials suggests that formulating curcumin improves systemic exposure and increases area under curve (AUC) and maximum blood concentration of curcumin, and thus, increases the bioavailability of curcumin ($C_{\text{max}}$) [41–44, 49, 50].
### Table 1  Composition of different curcumin formulations

| Formulation name                  | Technology          | Curcuminoid concentration | Ingredients                                                                 |
|-----------------------------------|---------------------|---------------------------|-----------------------------------------------------------------------------|
| Theracurmin [41]                  | Colloidal nanoparticles | 12%                       | 12% curcuminoids, 46% glycerine, 4% gum ghatti, 38% water                   |
| Meriva® [42]                      | Phytosome            | 18–20%                    | Curcumin, soy lecithin, microcrystalline cellulose                          |
| NovaSol [43]                      | Liquid micelles      | 6%                        | 93% Tween 80, and 7% curcumin powder                                        |
| CurQfen [49]                      | Soluble fibre blend  | Not defined               | Fenugreek soluble fibre blend, and 40% curcumin                             |
| Longvida® [44]                    | Solid lipid curcumin particle | 20–30%                  | Solid lipid curcumin particle lipids, phosphatidylcholine, and 20% curcumin |
| Curcumin C3 Complex® + Bioperine [50] | Not applicable     | Not defined               | Bioperine and curcuminoids                                                  |

### Table 2  Pharmacokinetic parameters of curcumin from the different curcumin-based formulations and reference (unformulated curcumin) in human studies

| Formulation                  | Clinical study design | Population                          | Intervention     | Dose                          | Sample hydrolysis | C<sub>max</sub> (ng/mL) | T<sub>max</sub> (h) | AUC<sub>0→t</sub> (ng h/mL) |
|-----------------------------|-----------------------|-------------------------------------|------------------|-------------------------------|-------------------|-------------------------|----------------|--------------------------|
| Theracurmin [41]            | Randomised, crossover | Asian 14 (8 males, 6 females) 44.1 ± 8.5 years | Formulation<sup>a</sup> | 30 mg theracurmin            | Hydrolysed        | 29.5 ± 12.9             | 1              | 113 ± 61<sup>†</sup>     |
|                            |                       |                                     | Control<sup>a</sup> | 30 mg curcumin powder         |                   | 1.8 ± 2.0               | 6              | 4.1 ± 7.0<sup>†</sup>    |
|                            |                       |                                     | Formulation<sup>b</sup> | 376 mg curcumin               | Hydrolysed        | 206.9 ± 54.9            | 2.7 ± 0.3     | 1336.0 ± 357.1<sup>†</sup> |
|                            |                       |                                     | Control<sup>b</sup> | 1799 mg curcumin              |                   | 14.4 ± 4.2              | 6.9 ± 2.2     | 202.8 ± 53.8<sup>†</sup> |
|                            |                       |                                     | Formulation<sup>a</sup> | 500 mg curcuminoids           | Hydrolysed        | 1189.1 ± 518.7          | 1.1 ± 0.4     | 4474.7 ± 1675.2<sup>‡</sup> |
|                            |                       |                                     | Control<sup>a</sup> | 500 mg curcuminoids           |                   | 2.6 ± 4.9               | 7.5 ± 8.2     | 24.1 ± 42.6<sup>‡</sup>  |
|                            |                       |                                     | Formulation<sup>a</sup> | 600 mg curcumin               | Not hydrolysed    | 0.4 ± 0.2 (µg/g)        | 1             | 8100 ± 287<sup>†</sup> (µg h/g) |
|                            |                       |                                     | Control<sup>a</sup> | 1000 mg curcumin              |                   | 0.02 ± 0.01 (µg/g)      | 0.5           | 510 ± 123<sup>†</sup> (µg h/g) |
|                            |                       |                                     | Formulation<sup>b</sup> | 650 mg curcuminoids           | Not hydrolysed    | 22.4 ± 1.9              | 2.4 ± 0.4     | 95.3 ± 4.6<sup>†</sup>   |
|                            |                       |                                     | Control<sup>b</sup> | 650 mg curcuminoids           |                   | < 1                     | ND            | ND                       |
|                            |                       |                                     | Formulation<sup>b</sup> | 2000 mg curcumin with bioperine | Not hydrolysed    | 180 ± 30                | 0.69 ± 0.07   | 80 ± 10<sup>†</sup>        |
|                            |                       |                                     | Control<sup>b</sup> | 2000 mg curcumin              |                   | 6 ± 5                   | 1             | 4<sup>†</sup>             |

<sup>a</sup> Mean ± standard deviation  
<sup>b</sup> Mean ± standard error of mean  
<sup>†</sup> AUC<sub>0–24</sub>  
<sup>‡</sup> AUC<sub>0–12</sub>  
<sup>†</sup> AUC<sub>0–6</sub>  

Control: unformulated curcumin, AUC area under the drug concentration–time curve, C<sub>max</sub> maximum drug concentration, T<sub>max</sub> time at maximum drug concentration, ND not defined.
emulsifier) that could deliver most of the curcumin to the intestinal wall for absorption by escaping the phase separation in the gastrointestinal tract [43]. Schiborr et al. [43] also observed that less than 0.2% of the oral dose of Novasol curcumin was excreted in urine within 24 h, and concluded that the remaining >98.8% of the ingested curcumin was either excreted via the bile and faeces or may have been distributed to body tissues where it may potentially exert biological activities. Meriva [42], a formulation using natural curcuminoids and lecithin (phosphatidylcholine phytosome complex of soy) in the ratio of 2:1 along with two parts of microcrystalline cellulose yielded the highest $T_{\text{max}}$ at 2.7 ± 1 h for a dose of 376 mg of curcuminoids, and 29-fold higher curcuminoid absorption compared to the unformulated curcumin. Interestingly, the authors [42] concluded that lecithin favoured the bioavailability of demethoxycurcumin as its plasma content was found to be higher than curcumin itself, despite its low concentration in the formulation.

The formulation of fenugreek fibre and 40% curcumin called CurQfen [51] yielded the highest AUC over 24 h at a dose of 1500 mg (equivalent to 600 mg curcumin). The use of soluble fibre in the formulation produced a non-digestible gel hydrocolloid that could ferment in the colon, prevent curcumin degradation in the gastrointestinal tract, and retard curcumin release resulting in a lag time of more than 5 h and less than 30% the total release after 24 h [51]. Thus, the fibre–curcumin complex contributed to improved and delayed curcumin absorption [51].

The curcumin formulation named Longvida [44] incorporated solid lipid curcumin particles (SLCP; 650 mg) in their formulation. A single dose of 130–195 mg of curcumin showed a $T_{\text{max}}$ of 2.4 h. The SLCP is a proprietary formula and comprises of curcumin mixed with soy lecithin containing purified phospholipids, docosahexaenoic acid (DHA), and/or vegetable stearic acid, ascorbyl (vitamin C) esters, and inert ingredients. The improved bioavailability of SLCP compared to unformulated curcumin is linked to key parameters such as curcumin/lipid/antioxidant ratio, globule-size distribution, and stability [51].

Consumption of a combination of curcumin and piperine in the Curcumin C3 Complex + Bioperine resulted in a 20-fold increase in plasma curcumin concentrations compared to the control formulation with the lowest $T_{\text{max}}$ of 0.69 h. Piperine is a P-glycoprotein and a uridine diphosphate-glucuronosyltransferase (UGT) inhibitor, and is suggested to improve absorption of curcumin by decreasing the efflux in the intestine and increasing the freely available curcumin in the systemic circulation [50]. Although curcumin was not observed in the plasma from 3 to 6 h, the bioavailability improved by 1.5 times compared to that of unformulated curcumin [50].

### Plasma curcumin concentration in exercise trials

Several studies [22, 23, 25, 52, 53] that investigated the effect of curcumin on exercise-induced inflammatory and oxidative stress markers also evaluated plasma curcumin concentration post-supplementation. All studies [22, 23, 25, 52, 53] observed an increase in plasma curcumin concentration post-supplementation at time points ranging from 2 h to 1–4 days (Table 3). The studies also concluded that supplementation with curcumin resulted in an increase in oxidative capacity [25], improvements in visual analogue scale for muscle soreness [23] and daily analysis of life demands questionnaire [52], and a decrease in CK levels [22, 53] (Table 4).

It is difficult to compare and evaluate the plasma curcumin concentration as studies by Tanabe et al. [22, 53] did not describe the sample preparation process and studies by Takahashi et al. [25], Sciberras et al. [52], and Tanabe et al. [23] hydrolysed their plasma samples and, therefore, the amount of curcumin in the plasma [48], may have been overestimated.

### Difficulties in comparing the effects of different pharmacokinetic characteristics of curcumin supplements on muscle damage markers

It is challenging to understand how the measured differences in pharmacokinetic characteristics ($C_{\text{max}}$, $T_{\text{max}}$, and AUC) of the specific supplements may relate to changes in inflammatory markers and attenuation of muscle damage. One key contributing factor is that although pharmacokinetic and exercise trials have both been carried out using the same formulations (Theracurmin [41] and Meriva [42]) different doses have been used for the exercise [23, 25, 43] versus the pharmacokinetic studies [41, 42]. In addition, some researchers have not measured the plasma curcumin concentrations post-supplementation, or do not clearly state the curcumin concentrations in the supplement, also making it challenging to compare results [24]. Others have hydrolysed the plasma samples before analysis [23, 25, 43], thus overestimating the true concentration of curcumin in the blood and making it difficult to accurately correlate any observed changes in levels of inflammatory markers to plasma curcumin levels. Finally, although some researchers [54] have reported that curcumin supplementation increased working capacity at the fatigue threshold and delayed the onset of neuromuscular fatigue, they did not analyse common inflammatory markers such as IL-6, TNF-α, and CK, also hindering comparisons between studies [54].
| Author and year | Population | Duration | Curcumin supplement | Dosage | Plasma curcumin concentration | Sample hydrolysis |
|----------------|------------|----------|---------------------|--------|------------------------------|-----------------|
| Takahashi et al., 2013 [25] | 26 males (26.8 ± 2.0 years) (recreationally active) | 1 day | Theracurmin | Control | Plasma curcumin concentrations in the double curcumin supplementation trial 2 h after exercise were significantly higher than those in the single curcumin supplementation trial | Hydrolysis |
| | | | | 90 mg/day 2 h before exercise 180 mg/day 2 h before and immediately after exercise |
| Tanabe et al., 2018 [22] | 10 males (28.5 ± 3.4 years) (untrained) | 7 days | Theracurmin | Experiment 1—180 mg (90 mg twice a day— at breakfast and dinner) consumed for 7 days before exercise | Plasma curcumin concentrations significantly decreased from baseline (38.8 ± 17.8 ng mL⁻¹) to 1, 3, 5, and 7 days after exercise (10.2 ± 5.4, 5.0 ± 2.7, 0.1 ± 0.4, and 0.1 ± 0.2 ng mL⁻¹, respectively) | No mention of the process used |
| | | | | | |
| | 10 males (29.0 ± 3.9 years) (untrained) | | Experiment 2—180 mg (90 mg twice a day— at breakfast and dinner) Consumed for 7 days after exercise | Plasma curcumin significantly increased from baseline (5.3 ± 6.1 ng mL⁻¹) to 1 day (50.6 ± 25.6 ng mL⁻¹), and then maintained a high plasma curcumin concentration through 3, 5, and 7 days after exercise (58.5 ± 37.1, 42.2 ± 46.0, and 41.1 ± 29.2 ng mL⁻¹, respectively) | |
| Tanabe et al., 2019 [23] | 8 males (28.0 ± 3.2 years) (untrained) | 7 days | Theracurmin CR-033P | Control | – | Hydrolysis |
| | | | | PRE—180 mg (90 mg twice a day— at breakfast and dinner) Consumed for 7 days before exercise | At baseline and at 1–3 d after exercise, the plasma curcumin concentration of the PRE group was significantly higher than that in the control and POST groups |
| | 8 males (28.8 ± 3.6 years) (untrained) | | | | |
| | 8 males (29.8 ± 3.4 years) (untrained) | | POST—180 mg (90 mg twice a day— at breakfast and dinner) Consumed for 4 days after exercise | At 1–4 d after exercise, the plasma curcumin concentration of the POST group was significantly higher than that of the control and PRE groups | |
Effect of curcumin on exercise-induced muscle damage

Intense training can lead to EIMD and can cause swelling, reduced ROM, and loss of muscle strength in the affected limb [16–18]. EIMD is characterised by muscular ultrastructural disruption that increases the release of inflammatory cytokines from myofibers and consequently increases their circulating levels. Muscle soreness increases from about 24 to 48 h post-exercise and decreases gradually from 72 h post-exercise. The activity of CK, a marker of EIMD, increases from 24 h onwards post-exercise and is sustained over a period of 7 days post-exercise [30].

Curcumin ingestion results in the attenuation of the release of inflammatory and oxidative markers, muscle pain, muscle performance, and CK levels by modulating inflammatory signalling cascades. Table 4 contains information from studies investigating the effect of curcumin on EIMD. Out of the 15 studies discussed below, the majority of participants in the trials were young physically active males 20–40 years of age. In addition, studies investigating the effect of curcumin supplementation on EIMD employed exercise protocols that led to different levels of muscle damage and assessed a variety of parameters, thus making it difficult to directly compare results and provide definitive conclusions as to whether curcumin supplementation is effective.

Muscle soreness

EIMD leads to delayed onset muscle soreness, associated with muscle pain, resulting in reductions in muscle strength and function and impairing physical function for several days post-exercise [55]. Damage to skeletal muscle activates phospholipase A₂, which leads to the removal of arachidonic acid from the cell membrane [56]. Arachidonic acid is converted to prostaglandin G₂ (PGG₂) under the influence of cyclooxygenase (COX-1 and COX-2) and then to prostaglandin H₂ (a common precursor to all prostaglandins) [27]. Prostaglandins are pro-inflammatory and cause redness, swelling (due to increased membrane permeability), and pain at the site of muscle damage [28]. Curcumin down-regulates the expression of COX-2 and thus, decreases the release of prostaglandins [29] which in turn reduces muscle soreness [19–23].

Studies [19–23] have shown significantly lower levels of muscle soreness when curcumin was consumed approximately 24 h before or after exercise. Consumption of 150 mg of curcumin (Theracurmin) immediately post-exercise resulted in a lower visual analogue scale (VAS) score for perceived muscle soreness compared to placebo at 48 and 72 h after unaccustomed squat exercises in untrained males [19]. Similarly, consumption of 180 mg of Theracurmin
Table 4 Summary of studies examining the effect of curcumin intake on exercise-induced muscle damage (EIMD)

| Author and year                  | Population          | Study design                                      | Duration | Curcumin supplement | Dosage               | Curcuminoids content                              | Activity type                                                                 | Key findings                        |
|----------------------------------|---------------------|--------------------------------------------------|----------|---------------------|----------------------|------------------------------------------------|--------------------------------------------------------------------------------|------------------------------------|
| Takahashi et al., 2013 [25]      | 26 males (26.8 ± 2.0 years) (recreationally active) | Double-blind, placebo-controlled, counterbalanced crossover design | 1 day    | Theracurmin         | Placebo              | NA                                             | Walking or running at 65% VO₂ max on a treadmill for 60 min                  | ↑ GPX |  
|                                 |                     |                                                   |          |                     | 90 mg/day 2 h before exercise 180 mg/day 2 h before and immediately after exercise | 10% curcumin, 2% curcuminoids without curcumin | ↑ Plasma curcumin ↑ BAP ↑ GSH ↑ TBARS ↑ GSSH ↑ SOD |                     |
|                                 |                     |                                                   |          |                     |                      | ↑ Plasma curcumin ↑ BAP ↑ GSH ↑ TBARS ↑ GSSH ↑ SOD | ↑ Plasma curcumin ↑ BAP ↑ GSH ↑ TBARS ↑ GSSH ↑ SOD |                     |
| Nakhostin-Roohi et al., 2016 [19]| 10 males (25.0 ± 1.6 years) (untrained) | Randomised controlled trial—double-blind crossover | Single dose | Theracurmin         | 150 mg Immediately post-exercise | 10% curcumin, 2% curcuminoids without curcumin | Uncustomed squat exercises | ↓ CK |  
|                                 |                     |                                                   |          |                     |                      |                                                   | ↓ Plasma curcumin ↓ IL-8            |                     |
| Tanabe et al., 2018 [22]        | 10 males (28.5 ± 3.4 years) (untrained) | Double-blind crossover design                     | 7 days   | Theracurmin         | 180 mg (90 mg twice a day—at breakfast and dinner) Consumed for 7 days before exercise | Not defined | 30 maximal eccentric contractions of the elbow flexors at an angular velocity of 120°/s |                     |
|                                 |                     |                                                   |          |                     | 180 mg (90 mg twice a day—at breakfast and dinner) Consumed for 7 days after exercise | Not defined | 30 maximal eccentric contractions of the elbow flexors at an angular velocity of 120°/s |                     |
|                                 | 10 males (29.0 ± 3.9 years) (untrained) |                                                   |          |                     | 180 mg (90 mg twice a day—at breakfast and dinner) Consumed for 7 days after exercise | Not defined | 30 maximal eccentric contractions of the elbow flexors at an angular velocity of 120°/s |                     |
| Tanabe et al., 2019 [23]        | 8 males (28.8 ± 3.6 years) (untrained) | Randomised, controlled, single-blind, parallel design study | 7 days   | Theracurmin CR-033P | PRE—180 mg (90 mg twice a day—at breakfast and dinner) Consumed for 7 days before exercise | 30% curcumin, 6% other curcuminoids | 30 maximal eccentric contractions of the elbow flexors at an angular velocity of 120°/s | ↑ Plasma curcumin at baseline |  
|                                 |                     |                                                   |          |                     | POST—180 mg (90 mg twice a day—at breakfast and dinner) Consumed for 4 days after exercise | 30% curcumin, 6% other curcuminoids | 30 maximal eccentric contractions of the elbow flexors at an angular velocity of 120°/s | ↑ Plasma curcumin at baseline |  
|                                 |                     |                                                   |          |                     |                      |                                                   |  
|                                 |                     |                                                   |          |                     |                      |                                                   | ↓ Plasma curcumin ↓ IL-8            |                     |

Key findings: ↑ = increased, ↓ = decreased, NA = not available

Activity type: Walking or running at 65% VO₂ max on a treadmill for 60 min

Key findings: ↑ GPX, ↑ d-ROM’s, ↑ Plasma curcumin, ↑ BAP, ↑ GSH, ↑ TBARS, ↑ GSSH, ↑ SOD

Key findings: ↑ Plasma curcumin, ↓ VAS for muscle soreness, ↓ CK
| Author and year | Population | Study design | Duration | Curcumin supplement | Dosage | Curcuminoids content | Activity type | Key findings |
|----------------|------------|--------------|----------|---------------------|--------|----------------------|--------------|-------------|
| Tanabe et al., 2015 [53] | 14 males (23.5 ± 2.3 years) (untrained) | Randomised, crossover design | Single dose | Theracurmin | 300 mg | Not defined | 50 maximal eccentric contractions of the elbow flexors at an angular velocity 120°/s | ↑ Plasma curcumin ↓ MVC torque ↓ CK ↑ IL-6 and TNF-α |
| McFarlin et al., 2016 [24] | 16 (5 M, 11 F) (20 ± 1 years) (untrained) | Randomised controlled | 6 days | Longvida | 400 mg | Not defined | 6 sets of 10 repetitions of the leg press exercise with a beginning load set at 110% of their estimated 1RM | ↑ Subjective quadriceps pain ↑ ADL ↑ CK ↓ TNF-α ↓ IL-8 ↑ IL-10 ↑ IL-6 |
| Sciberras et al., 2015 [52] | 11 males (35.5 ± 5.7 years) (trained) | Double-blind randomised crossover | 4 days | Meriva® | 500 mg with midday meal for 3 days and 500 mg just before exercise | Not defined | Participants exercised for 2 h at a power output equivalent to 95% of their lactate threshold | ↑ IL-6, IL-10 ↑ DALDA Questionnaire (better than usual) |
| F. Drobnic et al., 2014 [70] | C—9 males (32.7 ± 12.3 years) (trained) P—10 males (38.1 ± 11.1 years) (trained) | Randomised, placebo-controlled, single-centre, single-blind pilot trial | 4 days | Meriva® | 1 g twice a day 48 h prior to exercise and continued for 24 h after exercise | 200 mg/dose | Modified downhill running test—running at a constant speed for 45 min after a 10-min warm-up on treadmill | ↑ CRP ↑ hsCRP ↑ MCP-1 ↑ FRAP ↑ GPx ↑ CK ↓ IL-8 ↑ Intensity of pain |
| Jäger et al., 2019 [57] | 63 (31 M, 32 F) (21 ± 2 years) (trained) | Double-blind, randomised, placebo-controlled parallel design | 8 weeks | CurcuWIN® | Low-dose group 250 mg thrice a day (breakfast/lunch/dinner) High-dose group 1000 mg thrice a day (breakfast/lunch/dinner) | 50 mg of curcuminoids 200 mg of curcuminoids | 45-min downhill run at a ~ 15% grade and speed equivalent to 65% VO2Max after 5-min warm up | ↑ Subjective muscle pain (anterior, posterior) all groups ↑ Subjective (total) muscle pain in high-dose group 1 h and 24 h post-exercise ↑ Maximum bending torque and bending power in low-dose group |
Table 4 (continued)

| Author and year          | Population | Study design                       | Duration | Curcumin supplement | Dosage | Curcuminoids content | Activity type                                  | Key findings |
|--------------------------|------------|------------------------------------|----------|---------------------|--------|----------------------|-----------------------------------------------|--------------|
| McAllister et al., 2020  | 14 males   | Double-blinded, randomised, crossover design | 4 days   | CurcuFresh          | 1500 mg/day 1000 mg at breakfast and 500 mg at dinner for days 1, 2 and 3 and 45 min before testing on day 4 | 69 mg of curcuminoids | Dual stress challenges task that consisted of 35 min of steady-state exercise at a workload corresponding to 60% VO2 peak with mental stress challenges | † GHS ↓ SOD ↓ AOPP ↓ H2O2 |
|                          | (21–30 years) (trained) |                                  |          |                     |        |                      |                                               |              |
| S. Basham et al., 2019   | 20 males   | Randomised, double-blinded, placebo-controlled, crossover | 28 days  | CurcuFresh          | 1500 mg/day 1000 mg at breakfast and 500 mg at dinner for 28 days | 69 mg of curcuminoids | 225 repetitions of sit and stand using an aerobic step bench in 15 min | ↓ CK ↓ VAS † TAC † MDA † TNF-α |
|                          | (21.7 ± 2.9 years) (trained) |                                  |          |                     |        |                      |                                               |              |
| Amalraj et al., 2020     | 30 (12 M, 18 F) (36 ± 11 years) (trained) | Randomised, placebo-controlled, double-blind | 4 days   | Cureit               | 500 mg/day (consumed on day 2, 3 and 4 of study) | Not defined | Downhill running for 45 min | ↑ CK ↓ VAS for muscle pain |
|                          |            |                                    |          |                     |        |                      |                                               |              |
| Delecroix et al., 2017   | 16 males   | Randomised, placebo-controlled, balanced crossover design | 4 days   | MGD nature          | 2 g curcumin + 20 mg Piperine consumed three times a day every 6 h between 8 am and 10 pm On exercise day—45 min before, immediately post- and 6 h post-exercise | Not defined | 25 repetitions over 25 m of one-leg jumps on an 8% downhill slope | † Isometric peak torque † Concentric peak torque † Jump performance † CK † Muscle soreness at any time point |
|                          | (20.7 ± 1.4 years) (trained) |                                  |          |                     |        |                      |                                               |              |
| Herrick et al., 2020     | 47 (25 M, 22 F) (21.0 ± 2.6 years) (untrained) G1: 18 (C+FEN) G2: 14 (FEN) G3: 15 (P) | Randomised, double-blind, placebo-controlled, parallel design | 28 days  | CurQfen             | 500 mg/day curcumin + 300 mg fenugreek dietary fibre (galactomannans) Consumed in the morning before eating | 190 mg of curcumin | Maximal graded exercise test on a cycle ergometer | ↑ Physical working capacity at the fatigue threshold ↑ delay the onset of neuromuscular fatigue ↑ VO2 peak ↑ Time to exhaustion |
A study on untrained males reported a significantly lower perceived muscle soreness VAS score for the upper arm and elbow joint at 3 and 4 days after consumption of 180 mg Theracurmin CR-033P (90 mg twice a day at breakfast and dinner) for 4 days after eccentric elbow flexion [23]. Intake of 500 mg/day (Cureit; consumed on days 2, 3, and 4 after exercise) by trained participants resulted in a decreased pain VAS score from 2.90 to 1.17 in the curcumin group compared to a smaller decrease in the placebo group [20]. Moreover, administration of 2500 mg twice daily of curcumin for 2.5 days before exercise, and then 2500 mg twice daily for 2.5 days after exercise, resulted in moderate to large effect size reductions in muscle pain during single-leg squat and gluteal stretch at 24 and 48 h [21].

Not all studies show clear benefits of curcumin supplementation in reducing muscle soreness. For example, one study implementing 56 days of supplementation with 200 mg of curcuminoids (1000 mg curcumin by CurcuWIN) in trained individuals indicated not statistically significant reductions in muscle soreness. Compared to the placebo and low-dose (50 mg) curcumin groups, the treatment group showed non-significant reductions of 26, 20, and 8% lower muscle soreness immediately, 24, and 48 h post-exercise (conducted on 57 ± 3 days), respectively. Further research would be required to determine if the non-significant recovery benefits from curcumin may have been due to the final supplementation dose being administered 24 h prior to the exercise and the lack of supplementation after exercise [57].

A wide range of curcumin doses were administered across these studies with differences in formulation, frequency, and timing of ingestion relative to exercise, along with the training status of the participants and the exercise protocol used that may have influenced the extent of muscle soreness post-completion of exercise. In addition, VAS for soreness is a subjective measure, varying from individual to individual, thus potentially influencing the results. Nonetheless, studies have reported lower levels of soreness when 180–2500 mg/day curcumin was consumed immediately after exercise and/or within at least 24 h before and/or after exercise [21, 23].

Muscle performance

NF-κB is the transcriptional control for myokines (cytokines synthesised and released during muscular contractions) that are involved in post-exercise inflammatory responses [58]. Overuse of joints contributes to high mechanical stress and generates bone and cartilage extracellular matrix fragments that are recognised by receptors expressed by innate immune cells. Cell activation mediated by this process stimulates the
activation of NF-κB, resulting in secretion of inflammatory cytokines such as IL-1 and TNF-α, which contribute to tissue damage [59]. Curcumin blocks the signalling pathway of NF-κB, reducing the inflammatory response, decreasing swelling, while improving joint mobility and stiffness (assessed via maximum voluntary contraction (MVC) force and ROM) [22, 23, 53].

Intake of 90 mg curcumin (Theracurmin) twice a day (breakfast and dinner) for 7 days in untrained males resulted in significant improvements in MVC torque and ROM compared to the placebo group 3–7 days after eccentric exercise [22]. In a similar study, consumption of 90 mg curcumin (TheracurminCR-033P) twice a day (breakfast and dinner) for 4 days after eccentric exercise resulted in improvements in ROM of the elbow joint at 3–4 days in untrained males compared to the placebo group. The increases in the degree of ROM coincided with improvements in muscle soreness, indicating that the two could be related. However, another study [23] involving 30 maximal eccentric contractions of elbow flexors showed no significant differences in MVC torque between curcumin and placebo groups at all time points [23]. Muscle regeneration begins on day 3 after exercise [60], and since the follow-up of the recovery period was only 4 days, this time period may have been insufficient to establish the effect of curcumin on MVC torque [23].

One study examining the effect of 50 mg of curcumin (in the form of 250 mg of CurcuWIN® or 200 mg of curcumin (in the form of 1000 mg of CurcuWIN®) over 56 days in physically active men and women reported that curcumin could prevent decreases in peak extension torque values observed at 1 and 24 h after muscle-damaging exercise (downhill running) [57]. However, changes in isokinetic peak and average flexion torque, peak extension and flexion power, peak and average peak torque failed to yield statistical significance at 1, 24, 48, and 72 h post-exercise. One of the limitations of the supplementation protocol used was conducting the exercise protocol after discontinuing curcumin supplementation and it is possible that continuing supplementation may have prevented a decrease in the performance measures [57].

Ingestion of 150 mg of curcumin (Theracurmin) immediately and 12 h after eccentric exercise in untrained males led to a decrease in MVC from baseline [53]. However, the decrease in MVC was significantly smaller (33.0 ± 8.0%) in the curcumin group than in the placebo group (40.0 ± 9.1%) immediately after exercise and 48–96 h after exercise. The smaller decrease in MVC indicates that intake of curcumin leads to a lower level of muscle damage than the placebo group. However, ROM significantly decreased through all time points from the baseline in the curcumin group and there was no significant interaction effect for the changes compared to the placebo groups [53]. This suggests that supplementation with 150 mg Theracurmin twice a day for only one day is inadequate to improve ROM post-exercise.

In a randomised crossover trial, elite rugby players, consuming 6 g of curcumin and 60 mg of piperine (MGD Nature, Brandéron, France) each day starting 48 h pre-exercise and continuing until 48 h post-exercise experienced a moderately smaller loss of mean power output during the 6-s sprint compared to the control group 24 h after exercise [61]. However, this result was counterbalanced by the absence of effect from curcumin supplementation on isometric peak torque, concentric peak torque, and jump performance at all time points. This was possibly because the EIMD protocol consisted of 25 repetitions over 25 m of one-leg jumps on an 8% downhill slope, which required a greater neuromuscular recruitment pattern compared to isokinetic and isometric tests [61]. Furthermore, a possible limitation of this study could be the quick muscle damage recovery kinetics (training adaptations) which could be a result of the population tested, i.e. elite rugby players [61].

Another study examining the effects of curcumin in combination with fenugreek soluble fibre (CUR + FEN) or fenugreek soluble fibre alone (FEN) on the physical working capacity at the fatigue threshold (PWCFT), peak oxygen consumption (VO₂ peak), and time to exhaustion (Tlim) on a graded exercise test in untrained subjects for 28 days [54], showed no effect of curcumin supplementation on VO₂ peak or Tlim. However, the PWCFT was greater after combined supplementation with curcumin and FEN compared with a placebo in ~20% of subjects [54].

Thus, based on the results obtained by Tanabe et al. [23], improvements in both MVC force and ROM in untrained males could be a result of consumption of 180 mg of curcumin in divided doses (two times a day) across 4–7 days post-eccentric exercise [23].

**Creatine kinase**

Muscle creatine kinase (CK-MM), a marker of EIMD, is one of the three isoforms of CK, and is present at places within the muscle fibre where ATP consumption is high [62]. Eccentric muscle contractions exceeding the muscle’s resistance result in perforations in the sarcolemma and damage to the sarcomeres, leading to increased membrane permeability [30, 62, 63], and release of CK into the interstitial fluid that then enters circulation via the lymphatic system [32]. In addition, the production of prostaglandins under the influence of COX-2 leads to vascular hyperpermeability, which could further aid release of CK [31].

Curcumin supplementation indirectly lowers plasma CK activity in several ways. First, curcumin in the blood offers a membrane-protective effect by altering the structure of the membrane [34] and improving membrane integrity, thus reducing CK release into the blood [64]. Second, curcumin...
can suppress the regulation of the COX-2 pathway, reducing prostaglandin release and hence influence vascular permeability, ultimately decreasing the intracellular–intravascular flow of CK [31]. Lastly, curcumin’s antioxidant properties can suppress the activity of ROS generated during muscle contractions that would ordinarily contribute to muscle damage via CK release [25, 65].

Several studies [19, 21, 22, 24, 26, 53] have observed significantly lower CK activity in the curcumin supplemented group at doses of 150–5000 mg/day in both trained [21] and untrained individuals [24, 53]. Single-dose investigations with an intake of 150 mg curcumin (Theracurmin) post-eccentric exercise showed a lower rise in CK activity immediately 0, 24, 48, and 72 h post-exercise compared to the placebo group [19]. In addition, CK activity was significantly lower in the curcumin group compared to the placebo group at 24 h [19]. Intake of 180 mg of curcumin (Theracurmin) for 7 days (90 mg twice a day, at breakfast and dinner) after exercise in untrained men also resulted in lower CK activity compared to the placebo group [22]. Consumption of 300 mg curcumin (Theracurmin) in divided doses (150 mg 1 h before exercise and 150 mg 12 h after exercise) showed a significantly smaller peak CK activity for the curcumin group (3398 ± 3562 IU/L) compared to placebo (7684 ± 8959 IU/L) and lower activity at 48, 72 and 96 h compared to baseline CK levels [53]. Similarly, 1500 mg/day (CurcuFresh) of curcumin resulted in significantly lower CK activity (199.62 U/L in the curcumin group compared to 287.03 U/L in the placebo group) [66]. Supplementation with 5000 mg/day of curcumin (Eurofins Scientific Inc.) for 5 days (5 capsules twice daily for 2.5 days before exercise, then 5 capsules twice daily for 2.5 days after exercise) in trained men showed a small reduction in CK activity at 24 and 48 h (−22 to 29%; ±21 to −22%) compared to the baseline values [21]. Lower CK activity in the curcumin groups across the studies may suggest that myofibril damage due to exercise was attenuated by curcumin ingestion [53].

Curcumin supplementation before and/or after exercise can decrease CK activity post-exercise. However, the magnitude of the reduction varies from study to study. Several factors may have contributed to the differing outcomes, including the training status of the participants, exercise protocol, duration of the study, the timing of ingestion of the curcumin supplement, and the formulation of the curcumin supplement itself (including treatments and other ingredients that affect the rate of absorption).

**Inflammatory markers**

Curcumin exerts anti-inflammatory actions by hindering the activation of NF-κB, suppressing the activation and phosphorylation of JAK/STAT proteins, and inhibiting MAPK signalling that contribute to the production of inflammatory markers such as TNF-α, IL-6, and IL-8 at the site of muscle damage [35].

TNF-α: The effect of curcumin supplementation on reducing TNF-α levels in the blood has been evaluated in several studies [21, 22, 24, 26, 53], with equivocal observations reported. No significant differences in plasma TNF-α levels were found between the curcumin and placebo trials when 150 mg [53] and 180 mg of curcumin [22] were consumed by untrained males following elbow flexor eccentric exercise [22, 53]. This could be because the exercise protocol involved small muscle mass and was short in duration, and, therefore, did not affect the inflammatory cytokine levels in the blood [67]. In addition, as TNF-α has a very short half-life (15–30 min), plasma concentrations do not always reflect those produced by myocytes [68]. Moreover, the TNF-α levels were measured in the blood and not the muscle tissue. As concentrations of inflammatory markers and oxidative stress markers after exercise are different between muscle tissue and blood [69], the observation on the effect of curcumin on TNF-α levels post-exercise may be limited. In the study by McFarlin et al. [24], untrained subjects completed 6 sets of 10 repetitions of leg press exercise and consumed 400 mg/day of curcumin 48 h before exercise, up until 72 h after exercise. They observed that TNF-α levels were significantly lower with curcumin at 1 day (−25%), 2 days (−23%), and 4 days (−23%) compared to placebo and concluded that a minimum dose of 400 mg curcumin could be effective in decreasing circulating levels of TNF-α [24].

However, studies involving 1500 mg/day of curcumin supplementation (69 mg of curcuminoids) for 28 days (CurcuFresh) [26] and 5000 mg/day for 5 days (Eurofins Scientific Inc.) [21] reported no significant decrease in plasma TNF-α levels after exercise in trained males. Curcumin supplementation in both the studies [21, 26] was likely ineffective because physically active individuals were recruited and the participants’ aerobic training status (150 min of moderate-intensity aerobic activity or 30 min of vigorous-intensity aerobic activity per week) offered the stimulus for adaptations that contributed to lower resting levels of TNF-α (1.2 pg/mL), thus negating any potential anti-inflammatory benefits of curcumin supplementation [26]. In addition, the authors proposed that no significant decrease in TNF-α levels was observed in the trained individuals because the exercise protocol they used (7 sets of 10 eccentric single-leg press repetitions) failed to cause sufficient muscle damage due to muscle adaptations from prior physical activity [21].

In most studies [21, 22, 53], the exercise protocol used led to only minor increases in plasma TNF-α concentrations and reported no significant differences in plasma TNF-α levels in curcumin groups compared to placebo groups. Nonetheless, as observed for untrained individuals, a minimum of 400 mg curcumin supplementation before and after eccentric
exercise may result in lower increases in TNF-α levels post-eccentric exercise [24].

**IL-6 and IL-8** The effect of curcumin supplementation on IL-6 and IL-8 levels before and after exercise in trained and untrained individuals has been evaluated in several studies [21, 22, 24, 53, 70]. No significant decrease in post-exercise IL-6 levels was observed in the curcumin group when compared to placebo groups when supplemented with 300 mg (Theracurmin) [53] or 400 mg of curcumin (Longvida), respectively [24]. However, the exercise protocol chosen (6 sets of 10 repetitions of the leg press exercise with a beginning load set at 110% of their estimated 1-repetition maximum) may not have been sufficient to increase the pro-inflammatory cytokines in the body due to the muscle mass involved and the short duration of the activity [24]. In addition, blood samples were not taken immediately after exercise but were taken at 24, 48, 72, and 96 h post-exercise; therefore, the researchers may have missed observing changes in IL-6 levels which typically increase from 8 to 12 h post-eccentric exercise, and return to baseline levels by 24 h post-exercise [53, 71]. In contrast, Nicol et al. [21] observed a decrease in IL-6 values in the curcumin supplemented group (5000 mg curcumin; Eurofins Scientific Inc.) in trained participants after 7 sets of 10 eccentric single-leg press repetitions on a leg press machine at 24 h relative to immediately post-exercise [21]. However, they also observed an increase in IL-6 levels (small standardised differences) immediately post-exercise (31%; ±29%) and again at 48-h post-exercise (32%; ±29%) relative to baseline, thus, making the overall effect of curcumin supplementation unclear [21].

Supplementation with 180 mg of curcumin for 7 days prior to exercise (30 maximal eccentric contractions of the elbow flexors) in untrained males was associated with significantly lower plasma IL-8 levels 12 h after exercise than the placebo group [22]. This decrease in plasma IL-8 levels was associated with high concentrations of curcumin in the blood during and after exercise that suppressed the exercise-induced inflammatory effect [22]. However, no significant differences were observed when curcumin was ingested after exercise [22]. Conversely, intake of 400 mg/day of curcumin (Longvida) (2 days before exercise and for 3 days after exercise) resulted in a significantly lower IL-8 concentrations at day 1 (−21%) and day 2 (−18%) post-exercise (6 sets of 10 repetitions of leg press) compared to placebo in untrained individuals [24]. Furthermore, intake of 1 g of curcumin supplement (Meriva) twice a day 2 days before exercise and for 1 day after exercise (modified downhill running) in trained individuals also resulted in a significantly lower increase in plasma IL-8 levels 2 h post-exercise [70]. Thus, although the effect of curcumin supplementation on IL-6 levels remains unclear, curcumin intake before and after exercise may lower serum IL-8 levels in both trained and untrained individuals.

**Oxidative markers**

Reactive oxygen species are produced by a variety of extracellular and intracellular agents such as electron leakage from the mitochondrial respiratory chain and NADPH oxidases [72]. Exercise can induce oxidative stress by increasing oxygen utilisation up to 200-fold in active muscles and contributing to excessive amounts of ROS [73, 74], which can damage DNA, proteins, and lipids [75, 76] and affect exercise performance [77].

Reactive oxygen species contribute to oxidative stress and maintain inflammation by promoting the activation of NF-κB. During a sustained inflammatory response, accumulation of neutrophils within tissues provides a growth medium for producing oxidative enzymes, cytokines, and chemokines [78–80]. Curcumin can suppress the activation of NF-κB and potentially lead to elevated antioxidant responses by activating NRF2 [36], which upregulates the synthesis of antioxidant proteins that protect against oxidative damage triggered by injury and inflammation [37]. Thus, activation of NRF2 can improve the total antioxidant capacity of the body and reduce the harmful effects of ROS [36]. In addition, the phenolic OH group of curcumin has the potential to act as a ROS scavenger and a quencher of the lipid peroxidative side chain, thus reducing the activity of lipid hydroperoxides [81].

Curcumin supplementation (Theracurmin) with 90 mg/day (2 h before endurance exercise) and 180 mg/day (90 mg 2 h before and immediately after endurance exercise) in healthy men attenuated exercise-induced increases in the serum concentrations of derivatives of reactive oxygen metabolites (d-ROMs) and serum biological antioxidant potential (BAP), and also reduced plasma glutathione levels (GSH) post-exercise [25]. However, there were no significant increase observed in superoxide dismutase (SOD), and glutathione peroxidase (GPx) concentrations immediately after and post-2 h of exercise compared to the pre-values in both single and double curcumin supplementation groups [25]. In addition, supplementation with 150 mg curcumin (Theracurmin) after squat exercises in untrained males resulted in improved total antioxidant capacity (TAC) at 24 and 48 h post-exercise compared to the placebo group [19].

In a study [22] where curcumin (180 mg; Theracurmin) was supplemented for 7 days before and after exercise in a double-blind crossover study, no statistically significant improvements in serum concentrations of d-ROMs and BAP were observed in the curcumin and placebo groups as well as between groups over time [17]. It is possible that the exercise protocol used in this study (30 maximal eccentric contractions of the elbow flexors at an angular velocity of 120°/s) was insufficient to result in significant oxidative stress [22]. Similarly, curcumin supplementation (1500 mg/day (1000 mg breakfast, 500 mg dinner); CurcuFresh) for
28 days in trained males showed no significant improvements in TAC post-exercise (225 repetitions of sit and stand using aerobic step bench over 15 min) compared to the placebo. According to the authors, the lack of significant changes in TAC may be because the assay did not quantify changes in enzymatic antioxidants and had poor sensitivity [26]. In a trial comparing the effects of curcumin (1 g, Meriva, 2x per day, 2 days before, and 1 day after exercise) versus placebo on catalase (CAT) and GPx levels in aerobically trained males who completed a 2 h downhill run, levels of both enzymes tended to increase 2 h after exercise and returned towards baseline values 24 h after exercise in both curcumin and placebo groups [70]. The recruitment of aerobically trained participants may have limited muscle damage with the exercise protocol used, thus possibly explaining limited changes in CAT and GPx activity [65].

Thus, results from two studies indicate that supplementation with 90–180 mg curcumin 2 h before exercise or immediately after exercise may improve the antioxidant capacity of the body [19, 25]. However, more quality research is needed to clarify the optimal dose.

Based on the studies discussed in this review, curcumin is beneficial in alleviating exercised-induced muscle damage. However, due to the vast differences in the formulations of curcumin supplements and study protocols, it is difficult to determine a single dose that would be effective in reducing various inflammatory markers and increase the antioxidant enzymes such as SOD and GPx. Table 5 summarises the timing and dose of curcumin supplementation required to improve muscle soreness and performance, inflammatory markers, and oxidative markers associated with EIMD in trained and untrained participants after eccentric and endurance exercise.

**Limitations of research and future directions**

The curcumin formulations discussed in this review have been developed with varying amounts of curcuminoids and different ingredients that impact their bioavailability, thus making it complicated to suggest a single optimal dose for reducing inflammation post-exercise. However, formulation-specific doses can be suggested based on the scientific research for each particular product.

Most of the studies discussed in this review recruited young healthy participants and there are no data available for the effect of curcumin on EIMD in older adults. Older adults are of special interest as they experience sarcopenia: generalised loss of skeletal muscle mass and muscle strength with age. As a result, older individuals have low muscle mass, low muscle strength, and an increased body fat percentage that contributes to chronic inflammation and oxidative stress compared to young individuals [82]. Moreover, the body composition of older individuals with sarcopenia is significantly altered compared to that of a young population. Therefore, the results from the studies based on the effect of curcumin on EIMD in young trained and untrained individuals should not be extrapolated to a sarcopenic older population. In addition, research on the chronic effects of curcumin consumption based on specific formulations, in association with an appropriate exercise protocol are required to evaluate their effects on recovery from EIMD.

**Conclusion**

Curcumin is a challenging ergogenic aid to study due to its poor bioavailability and poor metabolism in the intestine and liver. However, new formulations using nanoparticles [41], phytosomes [42], micelles [43], and phospholipid complexes [44] have been developed which demonstrate improved bioavailability and their effects on EIMD have been investigated in humans. Overall, curcumin supplementation is most effective in reducing EIMD when consumed by untrained individuals. Supplementation immediately after exercise and/or within at least 24 h before and/or after exercise is highly recommended. Though the optimum amount of curcumin required to decrease serum IL-6 levels is still unclear, supplementation with 400–1000 mg of curcumin 1–2 times a day could be considered to aid in improving muscle performance and lowering circulating IL-8 levels. To date, one study in untrained males has shown decreases in TNF-α levels post-exercise following consumption of 400 mg/day of curcumin for a period of 6 days [24]; however, more studies are needed to confirm this effect. In addition, curcumin supplementation in the range of 150–5000 mg/day has been effective in decreasing CK levels in both untrained and trained individuals when consumed pre- and/or post-exercise. To improve the antioxidant capacity of the body post-exercise, curcumin supplementation with 90–180 mg curcumin 2 h before exercise or immediately after exercise may be effective [19, 25].

Curcumin impacts the production of several exercise-induced inflammatory markers. However, the amount of curcumin required to elicit changes in these inflammatory markers vary significantly. In addition, as no two studies follow the same protocol along with the same curcumin formulation, it is challenging to conclude the amount of curcumin required to reduce EIMD.

**Author contributions** KN reviewed papers, wrote, and revised the article and AA, KRM, SJL and NCB suggested improvements and reviewed the article.
| EIMD markers and oxidative markers | Author | Training status of participants | Exercise protocol | Formulation | Curcuminoids content | Duration | Dosage | Timing of dose |
|-----------------------------------|--------|---------------------------------|-------------------|-------------|----------------------|----------|--------|----------------|
| Muscle soreness                   | Nakhostin-Roohi et al., 2016 [19] | Untrained           | Unaccustomed squat exercises | Theracurmin 10% curcumin, 2% curcuminoids without curcumin | Single dose 150 mg | Immediately post-exercise |
| Muscle performance                | Tanabe et al., 2018 [22]         | Untrained           | 30 maximal eccentric contractions of the elbow flexors at an angular velocity of 120°/s | Theracurmin Not defined | 7 days 180 mg | 90 mg twice a day at breakfast and dinner, consumed for 7 days after exercise |
|                                   | Tanabe et al., 2019 [23]         | Untrained           | 30 maximal eccentric contractions of the elbow flexors at an angular velocity of 120°/s | Theracurmin 30% curcumin, 6% other curcuminoids | 4 days 180 mg | 90 mg twice a day at breakfast and dinner, consumed for 4 days after exercise |
|                                   | Amalraj et al., 2020 [20]        | Trained             | Downhill running for 45 min | Cureit Not defined | 4 days 500 mg/day | Consumed on day 2, 3 and 4 of study |
|                                   | Nicol et al., 2015 [21]          | Trained             | 7 sets of 10 eccentric single-leg press repetitions on a leg press machine | Eurofins scientific Inc Not defined | 5 days 5000 mg/day | 5 capsules (2.5 g curcumin) twice daily for 2.5 days prior to exercise, then 5 capsules twice daily for 2.5 days after exercise |
| Muscle performance                | Tanabe et al., 2018 [22]         | Untrained           | 30 maximal eccentric contractions of the elbow flexors at an angular velocity of 120°/s | Theracurmin Not defined | 7 days 180 mg | 90 mg twice a day at breakfast and dinner, consumed for 7 days after exercise |
|                                   | Tanabe et al., 2019 [23]         | Untrained           | 30 maximal eccentric contractions of the elbow flexors at an angular velocity of 120°/s | Theracurmin 30% curcumin, 6% other curcuminoids | 4 days 180 mg | 90 mg twice a day at breakfast and dinner, consumed for 4 days after exercise |
|                                   | Tanabe et al., 2015 [53]         | Untrained           | 50 maximal eccentric contractions of the elbow flexors at an angular velocity 120°/s | Theracurmin Not defined | Single dose 300 mg | 150 mg 1 h before exercise and 150 mg 12 h after exercise |
| EIMD markers and oxidative markers | Author | Training status of participants | Exercise protocol | Formulation | Curcuminoids content | Duration | Dosage | Timing of dose |
|---|---|---|---|---|---|---|---|---|
| Creatine kinase | Nakhostin-Roohi et al., 2016 [19] | Untrained | Unaccustomed squat exercises | Theracurmin | 10% curcumin, 2% curcuminoids without curcumin | Single dose | 150 mg | Immediately post-exercise |
| | Tanabe et al., 2018 [22] | Untrained | 30 maximal eccentric contractions of the elbow flexors at an angular velocity of 120°/s | Theracurmin | Not defined | 7 days | 180 mg | 90 mg twice a day at breakfast and dinner, consumed for 7 days after exercise |
| | McFarlin et al., 2016 [24] | Untrained | 6 sets of 10 repetitions of the leg press exercise with a beginning load set at 110% of their estimated 1RM | Longvida | Not defined | 6 days | 400 mg | 48 h before exercise and for 72 h after |
| | Tanabe et al., 2015 [53] | Untrained | 50 maximal eccentric contractions of the elbow flexors at an angular velocity 120°/s | Theracurmin | Not defined | Single dose | 300 mg | 150 mg 1 h before exercise and 150 mg 12 h after exercise |
| | S. Basham et al., 2019 [26] | Trained | 225 repetitions of sit and stand using an aerobic step bench in 15 min | CurcuFresh | 69 mg of curcuminoids | 28 days | 1500 mg/day | 1000 mg at breakfast and 500 mg at dinner |
| | Nicol et al., 2015 [21] | Trained | 7 sets of 10 eccentric single-leg press repetitions on a leg press machine | Eurofins scientific Inc | Not defined | 5 days | 5000 mg/day | 5 capsules (2.5 g curcumin) twice daily for 2.5 days prior to exercise, then 5 capsules twice daily for 2.5 days after exercise |
| Tumour necrosis factor-α | McFarlin et al., 2016 [24] | Untrained | 6 sets of 10 repetitions of the leg press exercise with a beginning load set at 110% of their estimated 1RM | Longvida | Not defined | 6 days | 400 mg | 48 h before exercise and for 72 h after |
| EIMD markers and oxidative markers | Author | Training status of participants | Exercise protocol | Formulation | Curcuminoids content | Duration | Dosage | Timing of dose |
|-----------------------------------|--------|---------------------------------|-------------------|-------------|---------------------|----------|--------|---------------|
| Interleukin-6                     | Nicol et al., 2015 [21] | Trained                       | 7 sets of 10 eccentric single-leg press repetitions on a leg press machine | Eurofins scientific Inc | Not defined | 5 days   | 5000 mg/day | 5 capsules (2.5 g curcumin) twice daily for 2.5 days prior to exercise, then 5 capsules twice daily for 2.5 days after exercise |
| Interleukin-8                     | McFarlin et al., 2016 [24] | Untrained                      | 6 sets of 10 repetitions of the leg press exercise with a beginning load set at 110% of their estimated 1RM | Longvida | Not defined | 6 days   | 400 mg   | 48 h before exercise and for 72 h after |
|                                   | F. Drobnic et al., 2014 [70] | Trained                        | Modified downhill running test—running at a constant speed for 45 min after a 10-min warm-up on treadmill | Meriva® | 200 mg/dose | 4 days   | 1 g twice a day | 48 h prior to exercise and continued for 24 h after exercise |
| Biological antioxidant potential and glutathione | Takahashi et al., 2013 [25] | Recreationally active | Walking or running at 65% VO₂ max on a treadmill for 60 min | Theracurmin | 10% curcumin, 2% curcuminoids without curcumin | 1 day    | 180 mg/day | 2 h before and immediately after exercise |
References

1. Dei Cas M, Ghidoni R (2019) Dietary Curcumin: Correlation between Bioavailability and Health Potential. Nutrients 11(9):2147. https://doi.org/10.3390/nu11092147

2. Priyadarsini KI (2014) The chemistry of curcumin: from extraction to therapeutic agent. Molecules 19(12):20091–20112. https://doi.org/10.3390/molecules191220091

3. Panahi Y, Hosseini MS, Khalili N, Naimi E, Simental-Mendía LE, Majeed M, Sahbekar A (2016) Effects of curcumin on serum cytokine concentrations in subjects with metabolic syndrome: a post-hoc analysis of a randomized controlled trial. Biomed Pharmacother 82:578–582. https://doi.org/10.1016/j.biopharm.2016.05.037

4. Belcaro G, Cesarone MR, Dugall M, Pellegrini L, Ledda A, Grossi MG, Togni S, Appendino G (2010) Efficacy and safety of Meriva®, a curcumin-phosphatidylcholine complex, during extended administration in osteoarthrosis patients. Altern Med Rev 15(4):337–344

5. Henrotin Y, Priem F, Mobasheri A (2013) Curcumin: a new paradigm and therapeutic opportunity for the treatment of osteoarthritis: curcumin for osteoarthritis management. Springerplus 2(1):56. https://doi.org/10.1186/2193-1801-2-56

6. Liu Q, Luo WT, Sze SC, Tong Y (2009) Curcin inhibits cell proliferation of MDA-MB-231 and BT-483 breast cancer cells mediated by down-regulation of NFkappaB, cyclinD and MMP-1 transcription. Phytomedicine 16(10):916–922. https://doi.org/10.1016/j.phymed.2009.04.008

7. Wu L, Guo L, Liang Y, Liu X, Jiang L, Wang L (2015) Curcumin suppresses stem-like traits of lung cancer cells via inhibiting the JAK2/STAT3 signaling pathway. Oncol Rep 34(6):3311–3317. https://doi.org/10.3892/or.2015.4279

8. Prasad S, Tyagi AK, Aggarwal BB (2014) Recent developments in delivery, bioavailability, absorption and metabolism of curcumin: the golden pigment from golden spice. Cancer Res Treat 46(1):2–18. https://doi.org/10.4143/crt.2014.46.1.2

9. Asp S, Rohde T, Richter EA (1997) Impaired muscle glycogen resynthesis after a marathon is not caused by decreased muscle GLUT-4 content. J Appl Physiol (1985) 83(5):1482–1485. https://doi.org/10.1152/jappl.1997.83.5.1482

10. Asp S, Daugaard JR, Richter EA (1995) Eccentric exercise decreases glucose transporter GLUT4 protein in human skeletal muscle. J Physiol 482(PT 3):705–712. https://doi.org/10.1113/jphysiol.1995.sp020553

11. Burt DG, Lamb K, Nicholas C, Twist C (2014) Effects of exercise-induced muscle damage on resting metabolic rate, sub-maximal running and post-exercise oxygen consumption. Eur J Sport Sci 14(4):337–344. https://doi.org/10.1080/17461391.2013.783628

12. Gomes JH, Mendes RR, Franca CS, Da Silva-Grigolote ME, Pereira da Silva DR, Antonioli AR, de Oliveira ESAM, Quintans-Júnior LJ (2020) Acute leukocyte, muscle damage, and stress marker responses to high-intensity functional training. PLoS ONE 15(12):e0243276. https://doi.org/10.1371/journal.pone.0243276

13. Giovanelli N, Biasutti L, Salvadego D, Alemayehu HK, Grassi B, Lazzer S (2020) Changes in skeletal muscle oxidative capacity after a trail running race. Int J Sports Physiol Perform 15(2):278–284. https://doi.org/10.1123/ijssp.2018-0882

14. Giovanelli N, Lazzer S, Caucci S (2020) Muscle damage and inflammatory status markers after a 3-stage trail running race. J Sports Med Phys Fitness 60(11):1486–1492. https://doi.org/10.1007/s40279-020-01355-z

15. Bontemps B, Vercruyssen F, Gruet M, Louis J (2020) Downhill running: what are the effects and how can we adapt? A narrative review? Sports Med 50(12):2083–2110. https://doi.org/10.1007/s40279-020-01355-z

16. Byrne C, Twist C, Eston R (2004) Neuromuscular function after exercise-induced muscle damage: theoretical and applied implications. Sports Med 34(1):49–69. https://doi.org/10.2165/00003940-200401000-00005

17. Hyldahl RD, Hubal MJ (2014) Lengthening our perspective: morphological, cellular, and molecular responses to eccentric exercise. Muscle Nerve 49(2):155–170. https://doi.org/10.1002/mus.24077

18. Mackey AL, Kjaer M (2017) The breaking and making of healthy adult human skeletal muscle in vivo. Skelet Muscle 7(1):24. https://doi.org/10.1186/s13395-017-0142-x

19. Nakhostin-Roohi B, Nasirvand Moradlou A, Mahmoodi Hamidi S, Ghanivand B (2016) The effect of curcumin supplementation on selected markers of Delayed Onset Muscle Soreness (DOMS). Ann Appl Sport Sci 4 (2):25–31. https://doi.org/10.18869/acadpub.aassjournal.4.2.25

20. Amalraj A, Divya C, Gopi S (2020) The effects of bioavailable curcumin (cureit) on delayed onset muscle soreness induced by eccentric continuous exercise: a randomized, placebo-controlled double-blind clinical study. J Med Food 23(5):545–553. https://doi.org/10.1089/jmf.2019.4533

21. Nicol LM, Rowlands DS, Fazakerly R, Kellett J (2015) Curcumin supplementation likely attenuates delayed onset muscle soreness (DOMS). Eur J Appl Physiol 115(8):1769–1777. https://doi.org/10.1007/s00421-015-3152-6

22. Tanabe Y, Chino K, Ohnishi T, Ozawa H, Sagayama H, Maeda S, Takahashi H (2019) Effects of oral curcumin ingested before or after eccentric exercise on markers of muscle damage and
hours of endurance cycling. J Int Soc Sports Nutr 12(1):5. https://doi.org/10.1186/s12970-014-0066-3

53. Tanabe Y, Maeda S, Akazawa N, Zempo-Miyaki A, Choi Y, Ra SG, Imazumi A, Otsuka Y, Nosaka K (2015) Attenuation of indirect markers of eccentric exercise-induced muscle damage by curcumin. Eur J Appl Physiol 115(9):1949–1957. https://doi.org/10.1007/s00421-015-1370-4

54. Herrick LP, Goh J, Menke W, Campbell MS, Fleenor BS, Abel MG, Bergstrom HC (2020) Effects of curcumin and fenugreek soluble fiber on the physical working capacity at the fatigue threshold, peak oxygen consumption, and time to exhaustion. J Strength Cond Res 34(12):3346–3355. https://doi.org/10.1519/JSC.0000000000003852

55. Järvinen TA, Järvinen M, Kalimo H (2013) Regeneration of injured skeletal muscle after the injury. Muscles Ligaments Tendons J 3(4):337–345

56. Vella L, Caldow MK, Larsen AE, Tassoni D, Dellá Gatta PA, Gran P, Russell AP, Cameron-Smith D (2012) Resistance exercise increases NF-κB activity in human skeletal muscle. Am J Physiol Regul Integr Comp Physiol 302(6):R667-673. https://doi.org/10.1152/ajpregu.00536.2011

57. Jäger R, Purpura M, Kerkscik CM (2019) Eight weeks of a high intensity interval training program decreases circulating cytokines and oxidative markers by curcumin supplementation in a physically active population: a systematic review. Nutrients 12(2):501. https://doi.org/10.3978/j.issn.2079-4926.2019.01.01

58. Herrick LP, Goh J, Menke W, Campbell MS, Bloomer RJ (2012) Effects of a fruit/berry/vegetable supplement on muscle function and oxidative stress in humans: cause and consequences. Free Radic Biol Med 51(5):942–950. https://doi.org/10.1016/j.freeradbiomed.2011.12.009

59. Powers SK, Nelson WB, Hudson MB (2011) Exercise-induced oxidative stress in humans: cause and consequences. Free Radic Biol Med 51(5):942–950. https://doi.org/10.1016/j.freeradbiomed.2011.12.009

60. Jones DA, Newham DJ, Round JM, Tolfree SE (1986) Experimental human muscle damage: morphological changes in relation to other indices of damage. J Physiol 375:435–448. https://doi.org/10.1113/jphysiol.1986.sp016126

61. Delecroix B, Abaida AE, Leduc C, Dawson B, Dupont G (2017) Curcumin and pipérine supplementation and recovery following exercise induced muscle damage: a randomized controlled trial. J Sports Sci Med 16(1):147–153

62. Brancaccio P, Maffulli N, Buonaruo R, Limongelli FM (2008) Serum enzyme monitoring in sports medicine. Clin Sports Med 27(1):1–18, vii. https://doi.org/10.1016/j.csm.2007.09.005

63. Brancaccio P, Maffulli N, Limongelli FM (2007) Creatine kinase monitoring in sport medicine. Br Med Bull 81–82:209–230. https://doi.org/10.1093/bmb/ldm014

64. Davis JM, Murphy EA, Carmichael MD, Zielinski MR, Groeschwitz CM, Brown AS, Gangemi JD, Ghaffar A, Mayer EP (2007) Curcumin effects on inflammation and performance recovery following eccentric exercise-induced muscle damage. Am J Physiol Regul Integr Comp Physiol 292(6):R2168-2173. https://doi.org/10.1152/ajpregu.00858.2006

65. Powers SK, Ji LL, Kazavias AN, Jackson MJ (2011) Reactive oxygen species: impact on skeletal muscle. Compr Physiol 1(2):941–969. https://doi.org/10.1002/cphy.c100054

66. McCullister MJ, Basham SA, Waldman HS, Smith JW, Butawan MB, Bloomer RJ (2020) Effects of curcumin on the oxidative stress response to a dual stress challenge in trained men. J Diet Suppl 17(3):261–272. https://doi.org/10.1080/19390211.2018.1515142

67. Hirose L, Nosaka K, Newton M, Laveder A, Kano M, Peake J, Suzuki K (2004) Changes in inflammatory mediators following eccentric exercise of the elbow flexors. Exerc Immunol Rev 10:75–90

68. Ma Y, Zhao S, Shen S, Fang S, Ye Z, Shi Z, Hong A (2015) A novel recombinant slow-release TNF-α-derived peptide effectively inhibits tumor growth and angiogenesis. Sci Rep 5:13595. https://doi.org/10.1038/srep13595

69. Rosendal L, Søgaard K, Jæger M, Sjøgaard G, Langberg H, Kristiansen J (2005) Increase in interstitial interleukin-6 of human skeletal muscle with repetitive low-force exercise. J Appl Physiol (1985) 98(2):477–481. https://doi.org/10.1152/japplphysiol.00130.2004

70. Drobnic F, Riera J, Appendino G, Togni S, Franceschi F, Valle X, Pons A, Tur J (2014) Reduction of delayed onset muscle soreness by a novel curcumin delivery system (Meriva®): a randomised, placebo-controlled trial. J Int Soc Sports Nutr 11(1):31. https://doi.org/10.1186/1550-2783-11-31

71. Miles MP, Andring JM, Pearson SD, Gordon LK, Kasper C, Depner CM, Kidd JR (2008) Diurnal variation, response to eccentric exercise, and association of inflammatory mediators with muscle damage variables. J Appl Physiol (1985) 104(2):451–458. https://doi.org/10.1152/japplphysiol.00572.2007

72. TavassoliFAR, Vodgani M, Salehi Z, Izad M (2020) The influence of reactive oxygen species in the immune system and pathogenesis of multiple sclerosis. Autoimmune Dis 2020:5793817. https://doi.org/10.1155/2020/5793817

73. Hattori N, Hayashi T, Nakachi K, Ichikawa H, Goto C, Tokudome Y, Kuriki K, Hoshino H, Shibata K, Yamada N, Tokudome M, Suzuki S, Nagaya T, Kobayashi M, Tokudome S (2009) Changes of ROS during a two-day ultra-marathon race. Int J Sports Med 30(6):426–429. https://doi.org/10.1055/s-0028-1121144

74. Powers SK, Nelson WB, Hudson MB (2011) Exercise-induced oxidative stress in humans: cause and consequences. Free Radic Biol Med 51(5):942–950. https://doi.org/10.1016/j.freeradbiomed.2011.12.009

75. Caputo F, Veggianti R, Ghilelli L (2012) Redox modulation of the DNA damage response. Biochem Pharmacol 84(10):1292–1306. https://doi.org/10.1016/j.bcp.2012.07.022

76. Goldfarb AH, Garten RS, Cho C, Chee PD, Chambers LA (2011) Effects of a fruit/berry/vegetable supplement on muscle function and oxidative stress. Med Sci Sports Exerc 43(3):501–508. https://doi.org/10.1249/MSS.0b013e3181f1ef48

77. Ferreira LF, Reid MB (2008) Muscle-derived ROS and thiol regulation in muscle fatigue. J Appl Physiol (1985) 104(3):853–860. https://doi.org/10.1152/japplphysiol.00953.2007

78. Schwane J, Johnson SR, Vandenbergker CB, Armstrong RB (1983) Delayed-onset muscular soreness and plasma CPK and LDH activities after downhill running. Med Sci Sports Exerc 15(1):51–56

79. Willoughby DS, McFarlin B, Bois C (2003) Interleukin-6 expression after repeated bouts of eccentric exercise. Int J Sports Med 24(1):15–21. https://doi.org/10.1055/s-2003-37197

80. Cannon JG, Orencole SF, Fielding RA, Meydani M, Meydani SN, Fiatarone MA, Blumberg JB, Evans WJ (1990) Acute phase response in exercise: interaction of age and vitamin E on neutrophils and muscle enzyme release. Am J Physiol 259(6 Pt 2):R1214-1219. https://doi.org/10.1152/ajpregu.1990.259.6.R1214

81. Singh U, Barik A, Singh BG, Priyadarsini KI (2011) Reactions of reactive oxygen species (ROS) with curcumin analogues: structure-activity relationship. Free Radic Res 45(3):317–325. https://doi.org/10.3109/10715762.2010.532493

82. Soysal P, Stubbs B, Luco D, Luchini C, Solmi M, Peluso R, Sergi G, Isik AT, Manzato E, Maggi S, Maggio M, Prina AM, Cosco TD, Wu Y-T, Veronese N (2016) Inflammation and frailty in the elderly: a systematic review and meta-analysis. Ageing Res Rev 31:1–8. https://doi.org/10.1016/j.arr.2016.08.006