

## **Appendix 1. Additional inclusion and non-inclusion criteria.**

<b>Inclusion criteria</b>
Age $\geq$ 40 years-old Written consent obtained Health insurance coverage
<b>Non-inclusion criteria</b>
History of symptomatic crystal or inflammatory arthritis Knee surgery $\leq$ 1 year Knee intra-articular injection of corticosteroids and/or hyaluronic acid $\leq$ 2 months Current use of intramuscular, intravenous or oral corticosteroids Uncontrolled diseases that may require intramuscular, intravenous or oral corticosteroids Knee trauma $\leq$ 2 months Neurologic disorders involving the lower limbs Inability to speak, write or read French language Participation in another biomedical research Contraindication to resveratrol or hypersensitivity to any of its constituents

## Appendix 2. Information about pre-specified outcomes

**WOMAC questionnaire.** The WOMAC questionnaire is a self-administered, disease-specific instrument validated for OA. It consists of 24 items grouped into 3 subscales: pain (5 questions), stiffness (2 questions) and physical function (17 questions), with higher scores indicating greater disease severity.

**OARSI-OMERACT response.** The OARSI-OMERACT response to intervention will be defined as an improvement in pain (assessed by an 11-point pain NRS) or in function (assessed by the WOMAC function subscore)  $\geq 50\%$  and absolute change  $\geq 20$ , or improvement in at least 2 of the 3 following: 1) pain  $\geq 20\%$  and absolute change  $\geq 10$ , 2) function  $\geq 20\%$  and absolute change  $\geq 10$ , 3) patient global assessment (assessed by an 11-point global assessment NRS)  $\geq 20\%$  and absolute change  $\geq 10$ .

**NRS:** numeric rating scale; **OARSI:** Osteoarthritis Research Society International; **OMERACT:** Outcome Measures in Rheumatology; **WOMAC:** Western Ontario and McMaster Universities Arthritis Index.

**Appendix 3. World Health Organisation–Uppsala Monitoring Centre causality categories (extract).**

<b>Causality term</b>	<b>Assessment criteria*</b>
<b>Certain</b>	<p>Event or laboratory test abnormality, with plausible time relationship to drug intake</p> <p>Cannot be explained by disease or other drugs</p> <p>Response to withdrawal plausible (pharmacologically, pathologically)</p> <p>Event definitive pharmacologically or phenomenologically (i.e. an objective and specific medical disorder or a recognized pharmacological phenomenon)</p> <p>Rechallenge satisfactory, if necessary</p>
<b>Probable/likely</b>	<p>Event or laboratory test abnormality, with reasonable time relationship to drug intake</p> <p>Unlikely to be attributed to disease or other drugs</p> <p>Response to withdrawal clinically reasonable</p> <p>Rechallenge not required</p>
<b>Possible</b>	<p>Event or laboratory test abnormality, with reasonable time relationship to drug intake</p> <p>Could also be explained by disease or other drugs</p> <p>Information on drug withdrawal may be lacking or unclear</p>
<b>Unlikely (not excluded)</b>	<p>Event or laboratory test abnormality, with a time to drug intake that makes a relationship improbable (but not impossible)</p> <p>Disease or other drugs provide plausible explanations</p>
<p>*All points should be reasonably complied with.</p>	

#### Appendix 4. Expected nature of a suspected adverse reaction.

Suspected adverse reaction
Dizziness
Epidymitis
Erythematous
Headache
Interactions with macrolides
Myalgia of the lower extremities
Nasopharyngitis
Nephrotoxicity was reported in <i>in vivo</i> animal studies
Rash
Somnolence