

### Review Form 3

Journal Name:	<a href="#">Journal of Advances in Medical and Pharmaceutical Sciences</a>
Manuscript Number:	Ms_JAMPS_129997
Title of the Manuscript:	Administration of Exosome-Based Treatment for Exogenous Ochronosis Caused by Prolonged Use of Topical Hydroquinone
Type of the Article	Case report

#### PART 1: Comments

	Reviewer's comment	Author's Feedback <i>(Please correct the manuscript and highlight that part in the manuscript. It is mandatory that authors should write his/her feedback here)</i>
<b>Please write a few sentences regarding the importance of this manuscript for the scientific community. A minimum of 3-4 sentences may be required for this part.</b>	<b>Exosome based treatment in dermatology and cosmetology is recently introduced and dermatologists have to be familiar with it. Types of exosomes, their uses, side effects, are important issues.</b>	Exosomes consist of plant-derived exosomes, human-derived exosomes, and lactobacillus-derived exosomes. In dermatology, exosome-based therapy is beneficial for stimulating skin regeneration and rejuvenation. Exosomes can inhibit melanogenesis, accelerate wound healing, act as anti-aging agents, and reduce inflammation. Potential side effects include pain, redness, swelling, and discomfort. Serious side effects, particularly with the administration of human-derived exosomes, include allergic reactions such as anaphylactic shock. Several case reports have also documented other side effects, such as skin necrosis and granuloma formation.
<b>Is the title of the article suitable? (If not please suggest an alternative title)</b>	yes	Thank you for the comments and suggestions provided.
<b>Is the abstract of the article comprehensive? Do you suggest the addition (or deletion) of some points in this section? Please write your suggestions here.</b>	yes	Thank you for the comments and suggestions provided.
<b>Is the manuscript scientifically, correct? Please write here.</b>	yes	This case report is prepared based on standard guidelines and writing principles.
<b>Are the references sufficient and recent? If you have suggestions of additional references, please mention them in the review form.</b>	Some are not appropriately cited in the manuscript, but they are recent and sufficient.	Thank you for the comments and suggestions provided.

**Review Form 3**

<p><b>Is the language/English quality of the article suitable for scholarly communications?</b></p>	<p>See my additional comments</p>	<p>Thank you for the comments and suggestions provided.</p>
<p><b>Optional/General</b> comments</p>	<p>Additional comments to the author:</p> <ol style="list-style-type: none"> <li>1. Page 2: replace “Exosome are” with “exosomes are”</li> <li>2. Page 2: replace “signalling” with “signaling”</li> <li>3. Page 2: replace “can be improved to reduce pigmentation and make it appear healthier” with “can be improved by reducing pigmentation and providing a healthier appearance”</li> <li>4. Page3: replace “many skincare products misuse topical hydroquinone for skin bleaching cream” with “many skincare products contain topical hydroquinone for skin bleaching”</li> <li>5. Page 4, line 2: reference (Brenner &amp; Hearing, 2008) is not appropriate. Replace it with reference (Bhattar et al., 2015)</li> <li>6. Page 4 line 6: replace reference (Brenner &amp; Hearing, 2008), with reference (Bhattar et al., 2015)</li> <li>7. Page 4: what can the topical application of exosomes add to the intradermal injection?</li> <li>8. Page 4: what is the benefit of the PDT Light, what color? (To maximize the delivery of exosomes to the dermal layer)? Where you have injected it???</li> <li>9. Page 4: within the 36days after the procedure, was there any topical treatment, or advice to avoid sun exposure?</li> <li>10. Page 4, what caused the telangiectasias of the patient? Has she used topical steroids?</li> <li>11. Page 4: what complications are to be expected?</li> </ol>	<p>Thank you for the comments and suggestions provided.</p> <p>Comment number 1-6 have been revised as requested.</p> <p>Comment number 7 : In addition to being administered intradermally to stimulate neocollagenesis, we also applied topical exosomes to increase collagen deposition, accelerate wound healing, and improve overall cosmesis, ensuring that the regeneration and neocollagenesis processes occur optimally (Ash et al., 2024) → <b>has been added in the case report as requested in page 5</b></p> <p>Comment number 8 : PDT light therapy using two types of light, each applied for 10 minutes: green light to repair damaged skin and soothe sensitive skin, and red light to enhance skin cell regeneration and improve the absorption of topical exosomes into the skin → <b>has been added in the case report as requested in page 5</b></p> <p>Intradermal exosome treatment is evenly injected across all areas of the face, including both cheeks, temples, forehead, nose, chin, and upper neck. → <b>has been added in the case report as requested in page 5</b></p> <p>Comment number 9 : After the first treatment, the patient received a topical regimen consisting of a skincare set that included a facial wash, toner, serum, and a morning cream containing niacinamide, alpha arbutin, 3-O-ethyl ascorbic acid, and tranexamic acid, as well as a night cream containing retinol. The patient also used a hybrid sunscreen with SPF 50 PA++++ → <b>has been added on page 2 in the case presentation section</b></p> <p>Comment number 10 : The patient has no history of topical steroid use. The patient only used bleaching cream containing hydroquinone for 8 months.</p> <p>Comment number 11 : Potential side effects include pain, redness, swelling, and discomfort. Serious side effects, particularly with the administration of human-derived exosomes, include allergic reactions such as anaphylactic shock. Several case reports have also documented other side effects, such as skin necrosis and granuloma formation → <b>The potential side effects have been explained on page 5</b></p>

**PART 2:**

	<p><b>Reviewer’s comment</b></p>	<p><b>Author’s comment</b> (if agreed with reviewer, correct the manuscript and highlight that part in the manuscript. It is mandatory that authors should write his/her feedback here)</p>
<p><b>Are there ethical issues in this manuscript?</b></p>	<p><i>(If yes, Kindly please write down the ethical issues here in details)</i></p>	