Manuscript Evaluation Form

PAPER TITLE	The GPCR Antagonistic Drug CM-20 Stimulates Mitochondrial Activity in Human RPE Cells
AUTHOR(S) NAME	Qing Chang, Siquan Chen and Tahua Yang

Sec. A: REFEREE'S ASSESSMENT	cross as approp	riate)				
Criterion	Excellen	nt		Good	Fair	Poor
Originality of the Topic	X					
Technical Quality				X		
Importance in its Field	portance in its Field x					
Style & Overall Representation				X		
Readily Understandable	X					
Suitability for the Journal	X					
Adequate Illustrations or Drawings					x	
English Language				X		
Description		Yes	No		Comments/Sug	gestions
Does the title represent manuscript's contents?	1	Х				
Is the Abstract accurate and concise?		Х				
Are the approaches/methods properly described?		Х				
Are the conclusions and interpretations sound?		Х				
Are the references properly cited?		Х				
Is this a new/original contribution?		х				
Is it within the scope of the journal?		х				
Is the sample size adequate for the study?		х				
Overall the Paper is Rated:	(Excelle 10 9	ent 8	7	6	5 4 3 2	Poor)

Sec. B: REFEREE'S RECOMMENDATION	NS		OTHER SPECIFIC CRITICISM	S
Accept with minor changes		Imperi	fect style	х
Accept with major changes	X	Too long		
Reject in current form, but may be resubmitted	d	Refere		
Reject, with no resubmission		Typog	raphical and Grammatical errors	х
PAPER TYPE: Research article	X Review a	rticle	Letter article	
Systematic review article	Case Rep	oort	Clinical Trial Study	
Other				

Have the following Reporting Guidelines been used by the authors in this study (if applicable)?

- Randomized Controlled Trials CONSORT Consolidated Standards of Reporting Trials
- **Observational Studies** STROBE Strengthening the Reporting of Observational studies in Epidemiology
- Systematic Review of Controlled Trials PRISMA Preferred Reporting Items for Systematic Reviews and Meta-Analyses
- Study of Diagnostic accuracy/assessment scale STARD Standards for the Reporting of Diagnostic Accuracy Studies
- Case Reports CARE for case reports

Yes	No
Yes	No

Does the manuscript comply for Ethical Guidelines (if applicable)?

- Has the author provided the approval from local institutional review board? Or does the manuscript conform to the Helsinki Declaration on human experiments in the "Methods" section?
- Yes No
- Has the approval of institutional ethics committee been obtained by the author regarding experimentation on animals or other species? Is the name of the species mentioned in the title abstract and "Methods" Section of the article?
- Yes No

• In case of experimentation on human subjects, has informed consent been taken?

Yes	No

Confidential Comments to the Editor (not for Transmission to Authors):

Review of the article entitled The GPCR Antagonistic Drug CM-20 Stimulates Mitochondrial Activity in Human RPE Cells, by Qing Chang, Siquan Chen and Tahua Yang

This article is a significant contribution, generally well written, within the scope of the journal and suitable for publication after some revision.

On page 1, in the objective of the abstract, CM-20 can be stated without brackets In conclusion of the abstract and in keywords, poly pharmacological and Poly pharmacology can be separated, as well as in the rest of the manuscript

On page 2, paragraph 1, line 7, use italic for *per se*

Line 14 and 15, the fifth sentence of this paragraph can be improved, use comma after AMD eyes, which is correlated

Paragraph 2, line 11, poly pharmacology and multi target can be separated

Lines 22 and 23, use abbreviation after the first mentioning and correct spelling for adrenal receptors (ARs) Paragraph 3, line 2, the same labeling should be used for CM-20 in the entire manuscript

Move heading 2 and subheading 2.1 to the next column

State the full term and the abbreviation at the first place of mentioning in the manuscript, in the rest of the text either the abbreviation or full term can be used

Paragraph 4, line 3, use one space between the number and the unite of measure in the entire manuscript, for example 37 °C

Line 5, state the full term before the abbreviation, 1 % fetal bovine serum (FBS)

State the full term and the abbreviation at the first place of mention in the manuscript, in the rest of the text either the abbreviation or full term can be used

Paragraph 5, lines 6 and 7, place the number and unite of measure in the same row, (12.5 µM)

Paragraph 6, lines 2 and 9, micro plate can be divided, as well as in the rest of the manuscript

Line 3, state the full term before the abbreviation, tetramethylrhodamine methyl ester (TMRM)

Lines 6 and 7, place (10 nM) in the same row

Paragraph 7, line 1, micro plate can be divided

On page 3, paragraph 2, line 6, include and present Figure 1A, as well as all other figures after their first mention within the text of the manuscript

At the end of the first sentence of the third paragraph, line 3, state (Fig. 2).

Paragraph 3, line 11 and paragraph 5, line 8, correct spelling no treated

Move subheading 3.3 to the next column

On page 4, in conclusion, line 9, poly pharmacological should be divided

Add to the list of abbreviations DMEM (Dulbecco's Modified Eagle Medium)

In the list of abbreviations, delete one space between the sign of equality and the appropriate term, as follows = retinal, = mitochondria, = G-protein, as well as TMRM =

Add issue numbers to references 2, 5, 9, 12, 16, 18, 20, 22 and 23.

Add last page numbers to references 2, 4, 5, 12 and 23.

Format references and the entire manuscript according to the style of the journal.

Page 6, in the legend of Fig. (4)., place (12.5 μ M, n=4) in the same row.

ments for the Authors (continue on another sheet, if necessary):

FIELD OF EXPERTISE OF REFEREE: Materials, chemistry, medicinal and pharmaceutical chemistry, chemical technologies, nanotechnologies, biomedical engineering, spectroscopy

Name & Affiliation of Referee: Tamara Jovanovic, Department of Biomedical Engineering, Faculty of Mechanical Engineering, University of Belgrade, Kraljice Marije 16, 11120 Belgrade, Serbia

Tamara Jovanovic, February 14, 2022

SIGNATURE OF REFEREE / DATE