

British Journal of Oral & Maxillofacial Surgery

Disorders of microcirculation in the mechanism of bisphosphonate osteonecrosis: preliminary study in rats --Manuscript Draft--

Manuscript Number:	BJOMS-D-19-00503R2
Article Type:	Full Length Article
Keywords:	Bisphosphonate osteonecrosis, zoledronate, sulodexide, dopplerography, doppler, Sulodexide as an osteonecrosis prevention drug.
Corresponding Author:	German Alexandrovich Kosach Pavlov First Saint Petersburg State Medical University St. Petersburg, RUSSIAN FEDERATION
First Author:	German Alexandrovich Kosach
Order of Authors:	German Alexandrovich Kosach Alexander Petrosyan Andrey Yaremenko, Prof. MD Anna Zubareva, Prof. MD Svetlana Kutukova, Docent Orazmurad Yagmurov, Prof. MD Svetlana Chefu, D.Bi.Sci. Victoria Molokova Veronica Ignatova Seraphim Kosach Timur Vlasov, Prof. MD
Manuscript Region of Origin:	RUSSIAN FEDERATION
Abstract:	<p>Purpose. Determine the possibilities of angioprotection and reduce osteonecrosis while using bisphosphonates.</p> <p>Material and methods. In our experiment, 27 rats were used that were divided into 3 groups: group 1 was injected with saline, group 2 was administered zoledronic acid for 6 weeks, group 3 was administered zoledronic acid for 6 weeks and the subsequent administration of sulodexide. After that there was modeling of extraction of the teeth. A linear blood flow velocity was studied using laser and high-frequency Doppler ultrasound in the periodontal area of an extracted tooth in rats with the application of vasoactive substance Acetylcholine 3% for 1 min. Evaluation of changes in the structure of bone tissue was carried out by computed tomography of the head of the rats studied, comparison of data with the control group and the saline group.</p> <p>Results. A rapid decrease in microcirculation was detected during the use of Zoledronic acid for 6 weeks. Lower decrease in microcirculation was detected after 3 week Sulodexide therapy during the use of Zoledronic acid for 6 weeks. A decrease in blood flow in the mucous membrane and, to a greater extent, in bone tissue was detected.</p> <p>Conclusion. Zoledronic acid causes a statistically significant impairment of the periosteal blood flow, mucous membrane due to a complex of disorders, which includes both the cellular component (impaired endothelium-dependent vasodilation of the mucous membrane vessels) and in reducing the microcirculation intensity in the mucous membrane and bone tissue. However, the use of sulodexide improves blood flow restoration and reduces the size of osteonecrosis that occurs.</p> <p>Bisphosphonate osteonecrosis, zoledronate, sulodexide, dopplerography Sulodexide as an osteonecrosis prevention drug.</p> <p>Keywords: Bisphosphonate osteonecrosis, zoledronate, sulodexide, dopplerography, Sulodexide as an osteonecrosis prevention drug.</p>



The British Journal of Oral & Maxillofacial Surgery
CONFIRMATION OF AUTHORSHIP

TITLE: Disorders of microcirculation in the mechanism
of bisphosphonate osteonecrosis: preliminary study in rats
REFERENCE NO: D-19-00503 RA

We, the undersigned, confirm that we are the joint authors of the above paper.

We confirm that all the authors have had material input into the submission.

We confirm that, to our knowledge, all the claims, statements and conclusions are true and are our jointly held opinions.

We confirm that we all accept the terms of publication of the publisher.

Signed:

Name: Geyman Kosach Signature: [Signature]

Name: Devonika Ignatova Signature: [Signature]

Name: Seraphim Kosach Signature: [Signature]

Name: Victoria Molokova Signature: [Signature]

Name: Svetlana Chefu Signature: [Signature]

Name: Timur Vlasov Signature: [Signature]

This form must be signed by all the authors of the above paper and the original scanned and emailed to the journal office (BJOMS@elsevier.com) or uploaded with the PDF of the manuscript.

If your paper is accepted, publication will not proceed until the fully completed form is received.



The British Journal of Oral & Maxillofacial Surgery
CONFIRMATION OF AUTHORSHIP

TITLE: Disorders of microcirculation in the mechanism of bisphosphonate osteonecrosis: preliminary study in rats
REFERENCE NO: D-19-00503R1

We, the undersigned, confirm that we are the joint authors of the above paper.

We confirm that all the authors have had material input into the submission.

We confirm that, to our knowledge, all the claims, statements and conclusions are true and are our jointly held opinions.

We confirm that we all accept the terms of publication of the publisher.

Signed:

Name: Alexander Petrosyan Signature: [Signature]

Name: Oyo Zmurod Yagmuruz Signature: [Signature]

Name: Svetlana Kutukova Signature: [Signature]

Name: Anna Zubayeva Signature: [Signature]

Name: Andrey Karyemko Signature: [Signature]

Name: Signature:

This form must be signed by all the authors of the above paper and the original scanned and emailed to the journal office (BJOMS@elsevier.com) or uploaded with the PDF of the manuscript.

If your paper is accepted, publication will not proceed until the fully completed form is received.