

## ABSTRACT&amp;REFERENCES

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## CESAREAN SECTION IN THE PERINATAL CENTER OF III LEVEL – INDICATIONS AND RISK FACTORS

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*In the structure of labour, the frequency of caesarean section should not exceed 15 %. Therefore, it is extremely important to clearly define clinical situations in which caesarean section can be abandoned and, conversely, identify pregnant women in whom it is advisable to plan a caesarean section to prevent urgent situations.*

**The aim of the study** was to study the structure of indications for emergency abdominal delivery in a level III medical department – the urban perinatal center of Kharkov to optimize childbirth tactics.

**Material and methods.** Clinical and statistical analysis of pregnancy and childbirth histories of 550 women in labour who gave birth in the Kharkov city perinatal center during 2018–2019 was performed. The structure of the indications and the caesarean section frequency was analysed depending on clinical and anamnestic data using descriptive statistics methods,  $\chi^2$  criterion and calculation of the odds ratio (OR) using the PSSP statistical software package.

**Results.** The highest OR values were in diabetes mellitus, burdened gynecological history and cardiovascular diseases (OR more than 5.0). Gestational hypertension, first labour, genital tract infections, nervous system diseases, preeclampsia, obesity of the digestive system diseases, large fetus and vegetative-vascular dystonia (OR from 2.108 to 4.113) had a lesser effect, myopia, first pregnancy, and late reproductive age (OR from 1.619 to 1.958).

**Conclusion.** The most common causes of emergency caesarean section were weak labour (29.5 %) and fetal distress (13.2 %). In 20.9 % of women – concomitant diseases of the mother and large fetus. Indications for emergency caesarean section most often occurred in women in labour with diabetes mellitus, weighed down by a gynecological history and with cardiovascular diseases

**Keywords:** complications of childbirth, caesarean section, indications, obstetric and gynecological history, extragenital pathology, risk factors, odds ratio

## References

- Joyce, A. M., Hamilton, B. E., Osterman, M. J. K. (2017). Births in the United States, 2016. NCHS Data Brief, 287, 1–8.
- Hure, A., Powers, J., Chojenta, C., Loxton, D. (2017). Rates and Predictors of Caesarean Section for First and Second Births: A Prospective Cohort of Australian Women. *Maternal and Child Health Journal*, 21 (5), 1175–1184. doi: <http://doi.org/10.1007/s10995-016-2216-5>
- Hobbs, A. J., Mannion, C. A., McDonald, S. W., Brockway, M., Tough, S. C. (2016). The impact of caesarean section on breastfeeding initiation, duration and difficulties in the first four months postpartum. *BMC Pregnancy and Childbirth*, 16 (1). doi: <http://doi.org/10.1186/s12884-016-0876-1>
- Li, H.-T., Luo, S., Trasande, L., Hellerstein, S., Kang, C., Li, J.-X. et al. (2017). Geographic Variations and Temporal Trends in Cesarean Delivery Rates in China, 2008–2014. *JAMA*, 317 (1), 69. doi: <http://doi.org/10.1001/jama.2016.18663>
- Moskviak-Lesniak, D. Ye., Krasivska, A. V., Zhygaliak, I. T. (2019). Research of the factors which increase level of caesarian section in 2017–2018. *Bulletin of Social Hygiene and Health Protection Organization of Ukraine*, 3 (81), 25–29. doi: <http://doi.org/10.11603/1681-2786.2019.3.10587>
- Tarasenko, K. V., Gromova, A. M., Shafarchuk, V. M., Nesterenko, L. A. (2019). The Increasing Frequency of Caesarean Section as a Problem of Modern Obstetrics. *Ukrainian Journal of Medicine, Biology and Sport*, 4 (5), 197–201. doi: <http://doi.org/10.26693/jmbs04.05.197>
- Caughey, A. B., Cahill, A. G., Guise, J. M., Rouse, D. J. (2014). Safe prevention of the primary cesarean delivery. *American Journal of Obstetrics and Gynecology*, 210 (3), 179–193. doi: <http://doi.org/10.1016/j.ajog.2014.01.026>
- Davey, M., Flood, M., Pollock, W., Cullinane, F., McDonald, S. (2019). Risk factors for severe postpartum haemorrhage: A population-based retrospective cohort study. *Australian and New Zealand Journal of Obstetrics and Gynaecology*. doi: <http://doi.org/10.1111/ajo.13099>
- Kim, J. H., Lee, S. M., Lee, Y. H. (2018). Risk factors for respiratory distress syndrome in full-term neonates. *Yeungnam University Journal of Medicine*, 35 (2), 187–191. doi: <http://doi.org/10.12701/yujm.2018.35.2.187>
- Rossi, A. C., Prefumo, F. (2019). Antepartum and intrapartum risk factors for neonatal hypoxic-ischemic encephalopathy. *Current Opinion in Obstetrics and Gynecology*, 31 (6), 410–417. doi: <http://doi.org/10.1097/gco.0000000000000581>
- Mavropulo, T. (2018). Kesariv rozlyn i stan zdorovia ditei. *Z turbotoiu pro Zhinku*, 2 (86), 10–14.
- Keag, O. E., Norman, J. E., Stock, S. J. (2018). Long-term risks and benefits associated with cesarean delivery for mother, baby, and subsequent pregnancies: Systematic review and meta-analysis. *PLOS Medicine*, 15 (1), e1002494. doi: <http://doi.org/10.1371/journal.pmed.1002494>
- Betran, A., Torloni, M., Zhang, J., Gülmezoglu, A. (2015). WHO Statement on Caesarean Section Rates. *BJOG: An International Journal of Obstetrics & Gynaecology*, 123 (5), 667–670. doi: <http://doi.org/10.1111/1471-0528.13526>
- Al-Qahtani, S., Heath, A., Quenby, S., Dawood, F., Floyd, R., Burdya, T., Wray, S. (2011). Diabetes is associated with impairment of uterine contractility and high Caesarean section rate. *Diabetologia*, 55 (2), 489–498. doi: <http://doi.org/10.1007/s00125-011-2371-6>

15. Danilack, V. A., Hutcheon, J. A., Triche, E. W., Dore, D. D., Muri, J. H., Phipps, M. G., Savitz, D. A. (2020). Development and Validation of a Risk Prediction Model for Cesarean Delivery After Labor Induction. *Journal of Women's Health*, 29 (5), 656–669. doi: <http://doi.org/10.1089/jwh.2019.7822>

16. Tolcher, M. C., Holbert, M. R., Weaver, A. L., McGree, M. E., Olson, J. E., El-Nashar, S. A. et. al. (2015). Predicting Cesarean Delivery After Induction of Labor Among Nulliparous Women at Term. *Obstetrics & Gynecology*, 126 (5), 1059–1068. doi: <http://doi.org/10.1097/aog.0000000000001083>

17. Guan, P., Tang, F., Sun, G., Ren, W. (2020). Prediction of emergency cesarean section by measurable maternal and fetal characteristics. *Journal of Investigative Medicine*, 68 (3), 799–806. doi: <http://doi.org/10.1136/jim-2019-001175>

18. Elnakib, S., Abdel-Tawab, N., Orbay, D., Hassanein, N. (2019). Medical and non-medical reasons for cesarean section delivery in Egypt: a hospital-based retrospective study. *BMC Pregnancy and Childbirth*, 19 (1). doi: <http://doi.org/10.1186/s12884-019-2558-2>

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#### EFFECTIVENESS OF ARTERIAL HYPERTENSION CONTROL USING AMBULATORY BLOOD PRESSURE MONITORING IN PATIENTS IN THE REMOTE PERIOD AFTER MYOCARDIAL INFARCTION AND ISCHEMIC STROKE

p. 8-14

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*Arterial hypertension is one of the most important modified risk factors and its adequate control is a cornerstone of primary and secondary prevention. On the other hand, excessive lowering of blood pressure can be harmful, especially in patients with a history of ischemic events.*

**The aim:** To evaluate daily blood pressure fluctuations with ambulatory blood pressure monitoring in patients with controlled hypertension according to office measurements in the remote period after myocardial infarction and ischemic stroke

**Materials and methods:** The study involved 64 patients with hypertension in remote period (at least 6 months) after myocardial infarction (38 patients) and ischemic stroke (26 patients) with systolic blood pressure during the last three months and at least two visits. in the range of 120-139 mm Hg and diastolic blood pressure 70–89 mm Hg. Patients with valvular heart disease, severe arrhythmias, significant heart failure (NYHA III–IV) were not included in the study. All patients underwent daily blood pressure monitor-

ing. According to the recommendations of ESC/ESH in 2018, the optimal blood pressure control was considered to be the average daily blood pressure after myocardial infarction 130–140/70–79 mm Hg. at the age of >65 years, 120–130/70–79 mm Hg at the age of ≤65 years; after ischemic stroke 120–130/<80 mm Hg. All values below the optimal levels were considered to be low blood pressure, higher than optimal – high blood pressure. Particular attention was paid to excessive reduction of blood pressure with the calculation of the number of excessive reductions and their duration during the day.

**Results:** The average daily blood pressure values corresponded to the definition of optimal in only 23.4 % of patients with ischemic events. In 63.2 % of patients after myocardial infarction, the mean daily values of systolic blood pressure and diastolic blood pressure were lower than recommended, and in patients after ischemic stroke, this number was significantly lower – 23.1 %. Conversely, in more than half of the patients after ischemic stroke (53.6 %) the average daily blood pressure levels exceeded the recommended ones. In patients after myocardial infarction, this number was only 13.2 %.

Excessive reductions in blood pressure were recorded in 58 of 64 patients (90.6 %), in 54 persons such episodes were observed during the day (84.3 %). Persistent (over 1 hour) excessive decrease in blood pressure was registered in 16 people (25.0 %), 8 of whom had persistent excessive blood pressure reductions ≥2/day (12.5 %). Predictors of excessive blood pressure lowering were males, myocardial infarction, beta-blockers.

**Conclusions:** Patients with hypertension after myocardial infarction or ischemic stroke require regular monitoring of blood pressure, even in cases where blood pressure is within the recommended range for some time. It is necessary to take into account the risk not only of increasing but also of excessive lowering of blood pressure

**Keywords:** arterial hypertension, ambulatory blood pressure monitoring, myocardial infarction, ischemic stroke, blood pressure, arterial hypotension

#### References

- Williams, B., Mancia, G., Spiering, W., Agabiti Rosei, E., Azizi, M., Burnier, M. et. al. (2018). 2018 ESC/ESH Guidelines for the management of arterial hypertension. *European Heart Journal*, 39 (33), 3021–3104. doi: <https://doi.org/10.1093/eurheartj/ehy339>
- Richards, A. M., Nicholls, M. G., Troughton, R. W., Lainchbury, J. G., Elliott, J., Frampton, C. et. al. (2002). Antecedent hypertension and heart failure after myocardial infarction. *Journal of the American College of Cardiology*, 39 (7), 1182–1188. doi: [https://doi.org/10.1016/s0735-1097\(02\)01737-0](https://doi.org/10.1016/s0735-1097(02)01737-0)
- D'Agostino, R. B., Belanger, A. J., Kannel, W. B., Cruickshank, J. M. (1991). Relation of low diastolic blood pressure to coronary heart disease death in presence of myocardial infarction: the Framingham Study. *BMJ*, 303 (6799), 385–389. doi: <https://doi.org/10.1136/bmj.303.6799.385>
- Flack, J. M., Neaton, J., Grimm, R., Shih, J., Cutler, J., Ensrud, K., MacMahon, S. (1995). Blood Pressure and Mortality Among Men With Prior Myocardial Infarction. *Circulation*, 92 (9), 2437–2445. doi: <https://doi.org/10.1161/01.cir.92.9.2437>
- Rothwell, P. M., Howard, S. C., Dolan, E., O'Brien, E., Dobson, J. E., Dahlöf, B. et. al. (2010). Prognostic significance of visit-to-visit variability, maximum systolic blood pressure, and episodic hypertension. *The Lancet*, 375 (9718), 895–905. doi: [https://doi.org/10.1016/s0140-6736\(10\)60308-x](https://doi.org/10.1016/s0140-6736(10)60308-x)
- Antonini, L., Colivicchi, F., Greco, S., Guido, V., Malfatti, S., Gandolfi, A. et. al. (2003). Ambulatory Blood Pressure

Monitoring After Acute Myocardial Infarction. High Blood Pressure & Cardiovascular Prevention, 10 (2), 75–80. doi: <https://doi.org/10.2165/00151642-200310020-00003>

7. Zhang, Y., Wang, H., Xu, K., Wang, P., Li, X.-Y., Zhao, J.-B., Tang, Y. (2018). Ambulatory blood pressure variability within the first 24 hours after admission and outcomes of acute ischemic stroke. Journal of the American Society of Hypertension, 12 (3), 195–203. doi: <https://doi.org/10.1016/j.jash.2017.12.012>

8. Shin, D. H., Song, S., Lee, Y. B. (2019). Comparison of the Effect of Fimasartan versus Valsartan on Blood Pressure Variability in Acute Ischemic Stroke: A Double-Blind Randomized Trial. Cardiovascular Therapeutics, 2019, 1–8. doi: <https://doi.org/10.1155/2019/7836527>

9. Georgiou, E., Georgiou, P. I., Petidis, K., Markakis, K., Zografou, I., Karagiannis, A. (2019). Effect of Nebivolol and Olmesartan on 24-Hour Brachial and Aortic Blood Pressure in the Acute Stage of Ischemic Stroke. International Journal of Hypertension, 2019, 1–9. doi: <https://doi.org/10.1155/2019/9830295>

10. Xie, X., Xu, J., Gu, H., Tao, Y., Chen, P., Wang, Y., Wang, Y. (2017). The J-curve Association between Systolic Blood Pressure and Clinical Outcomes in Ischemic Stroke or TIA: The BOSS Study. Scientific Reports, 7 (1). doi: <https://doi.org/10.1038/s41598-017-10887-w>

11. Chwojnicky, K., Yagansky, A., Wierucki, L., Sichkaruk, I., Sydor, N., Nyka, W. M. et. al. (2011). Assessment of the Quality of Secondary Ischemic Stroke Prevention in Selected Urban Areas of Poland and Ukraine: The ASCEP Study Results. European Neurology, 65 (6), 323–331. doi: <https://doi.org/10.1159/000327576>

12. Yahenskyi, A. V., Sichkaruk, I. M. (2019). Prykhylnist do likuvannya patsientiv u viddalenyi period pislia infarktu miokarda. Ratsionalna farmakoterapiya, 1-2, 24–27.

13. Sleight, P., Redon, J., Verdecchia, P., Mancia, G., Gao, P., Fagard, R. et. al. (2009). Prognostic value of blood pressure in patients with high vascular risk in the Ongoing Telmisartan Alone and in combination with Ramipril Global Endpoint Trial study. Journal of Hypertension, 27 (7), 1360–1369. doi: <https://doi.org/10.1097/hjh.0b013e32832d7370>

14. Mancia, G., Kjeldsen, S. E., Zappe, D. H., Holzhauer, B., Hua, T. A., Zanchetti, A. et. al. (2015). Cardiovascular outcomes at different on-treatment blood pressures in the hypertensive patients of the VALUE trial. European Heart Journal, 37 (12), 955–964. doi: <https://doi.org/10.1093/eurheartj/ehv633>

15. Anderson, R. J., Bahn, G. D., Moritz, T. E., Kaufman, D., Abaira, C., Duckworth, W. (2010). Blood Pressure and Cardiovascular Disease Risk in the Veterans Affairs Diabetes Trial. Diabetes Care, 34 (1), 34–38. doi: <https://doi.org/10.2337/dc10-1420>

16. Kotti, K., Bagarhatta, R., Rathore, M., Bagarhatta, P. (2018). Is ambulatory blood pressure measurement a new indicator for survival among advanced heart failure cases. Indian Heart Journal, 70, S73–S78. doi: <https://doi.org/10.1016/j.ihj.2017.08.028>

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#### EFFICIENCY OF DOUBLE FIXED COMBINED THERAPY IN PATIENTS WITH ARTERIAL HYPERTENSION AND ASSOCIATED THYROTOXICOSIS AFTER ONE YEAR OF TREATMENT

p. 15-21

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**The aim:** to evaluate the effectiveness of different fixed double antihypertensive combinations based on the results of outpatient monitoring of arterial pressure and to analyze their effect on central arterial pressure and cardiac remodeling in patients with arterial hypertension and thyrotoxicosis.

**Materials and methods:** patients with hypertension and TT who were included in the study were divided into groups using the blind envelope method with compensated TT: patients of group 1 were prescribed a fixed combination of perindopril with indapamide, group 2 – a combination of perindopril with amlodipine. Patients with uncompensated TT were prescribed a combination of bisoprolol with perindopril. The groups were statistically comparable in terms of age, duration of AH and TT, SBP and DBP levels. The observation of the patients lasted 12 months. After a year of follow-up, AMAD and EchoCG were performed in order to study AHT in patients with hypertension and TT. The patients were consulted by a cardiologist and endocrinologist.

**Results:** After 6 months of treatment, 8 (16 %) patients were excluded from the study, of which 6 (75 %) due to the need for surgical treatment and did not achieve euthyroidism, 2 (25 %) due to the need for antiarrhythmic therapy. Studied double fixed combinations effectively reduce blood pressure (BP) according to the results of ambulatory blood pressure monitoring (ABPM).

**Conclusions:** The fixed combination of perindopril with indapamide, perindopril with amlodipine, bisoprolol with perindopril provided a decrease in both blood pressure according to ABPM results and central pressure, however, this was not accompanied by a significant improvement in the structural and functional state of the heart

**Keywords:** cardiac remodeling, thyrotoxicosis, arterial hypertension, fixed combination antihypertensive therapy, ambulatory blood pressure monitoring

#### References

1. Basnet, S., Dhital, R., Tharu, B., Ghimire, S., Poudel, D. R. (2018). Association between thyroid abnormalities and hypertension among hospitalized US patients: data from the National Inpatient Sample. Cardiovascular Endocrinology & Metabolism, 7 (4), 97–98. doi: <http://doi.org/10.1097/xce.0000000000000155>

2. Yildiz, C., Altay, M., Yildiz, S., Çağır, Y., Akkan, T., Ünsal, Y. A., Beyan, E. (2019). Arterial stiffness in hyperthyroid patients is deteriorated due to thyroid hormones. Archives of Endocrinology and Metabolism, 63 (3), 258–264. doi: <http://doi.org/10.20945/2359-3997000000135>

3. Amosova, K. M., Shyshkina, N. V., Rokyta, O. I., Rudenko, Yu. V., Katsytadze, I. Yu. (2016). Comparative evaluation of changes in brachial and central blood pressure in different categories of patients with uncomplicated arterial hypertension and determinants of pulse pressure amplification. Sertse i sudyny, 1, 37–45.

4. Berta, E., Lengyel, I., Halmi, S., Zrínyi, M., Erdei, A., Harangi, M. et al. (2019). Hypertension in Thyroid Disorders. *Frontiers in Endocrinology*, 10. doi: <http://doi.org/10.3389/fendo.2019.00482>

5. Alekseeva, O. A., Shaposhnik, I. I., Bogdanov, D. V. (2019). Restoration of clinical and instrumental parameters of the cardiovascular system in patients with diffuse toxic goiter after resolution of thyrotoxicosis. *Almanac of Clinical Medicine*, 47 (2), 138–148. doi: <http://doi.org/10.18786/2072-0505-2019-47-013>

6. Kniaskova, I. I., Zhadan, A. V., Nesen, A. O. (2017). Arterialna ryhidnist yak chynnyk ryzyku i likuvalna mishen pry arterialnii hipertenzii. *Praktychna anhiolohiia*. 1 (76), 5–14.

7. Tsymbaliuk, I. L. (2016). Optyimizatsiia likuvannia ta profilaktyky sertsevo-sudynnykh porushen u khvorykh na arterialnu hipertenziiu, poiednanu z tyreotoksykozom. *Zaporizhzhia*.

8. Williams, B., Mancia, G., Spiering, W., Agabiti Rosei, E., Azizi, M., Burnier, M. et al. (2018). 2018 Practice guidelines for the management of arterial hypertension of the European Society of Cardiology and the European Society of Hypertension. *Blood Pressure*, 27 (6), 314–340. doi: <http://doi.org/10.1080/08037051.2018.1527177>

9. Ross, D. S., Burch, H. B., Cooper, D. S., Greenlee, M. C., Laurberg, P., Maia, A. L. et al. (2016). 2016 American Thyroid Association Guidelines for Diagnosis and Management of Hyperthyroidism and Other Causes of Thyrotoxicosis. *Thyroid*, 26 (10), 1343–1421. doi: <http://doi.org/10.1089/thy.2016.0229>

10. Marwick, T. H., Gillebert, T. C., Aurigemma, G., Chirinos, J., Derumeaux, G. (2017). Recommendations on the use of echocardiography in adult hypertension: a report from the European Association of Cardiovascular Imaging (EACVI) and the American Society of Echocardiography (ASE). *Systemic Hypertension*, 14 (2), 6–28. doi: [http://doi.org/10.26442/2075-082x\\_14.2.6-28](http://doi.org/10.26442/2075-082x_14.2.6-28)

11. Babenko, A. Yu. (2014). Cardiovascular disorders in thyrotoxicosis of various origins: optimization of treatment approaches. *Mezhdunarodnii endokrinologicheskii zhurnal*, 8 (64), 65–78.

12. Hegedus, L. (2018). Does therapy for thyroid dysfunction decrease mortality? *Endocrine Abstracts*. Barcelona. doi: <http://doi.org/10.1530/endoabs.56.p16>

13. Lillevang-Johansen, M., Abrahamsen, B., Jørgensen, H. L., Brix, T. H., Hegedüs, L. (2019). Duration of Hyperthyroidism and Lack of Sufficient Treatment Are Associated with Increased Cardiovascular Risk. *Thyroid*, 29 (3), 332–340. doi: <http://doi.org/10.1089/thy.2018.0320>

14. Cherenko, M. S. (2016). The current opinion on management and treatment of hyperthyroidism and other forms of thyrotoxicosis: review of the latest Guidelines of American Thyroid Association (2016). *Clinical Endocrinology and Endocrine Surgery*, 4 (56), 87–94. doi: [http://doi.org/10.24026/1818-1384.4\(56\).2016.87324](http://doi.org/10.24026/1818-1384.4(56).2016.87324)

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#### KERATITIS CAUSED BY *PSEUDOMONAS AERUGINOSA*: TREATMENT IN THE EXPERIMENT

p. 21-28

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*The aim of the study was to investigate the effectiveness of treatment of keratitis caused by *Pseudomonas aeruginosa* using official ophthalmic forms of antibiotics, which are effective against the pathogen.*

**Materials and methods.** Adult 36 rabbits weighing 3–3.5 kg were undergone the causing of purulent keratitis by applying a clinical strain of *P. aeruginosa* as a suspension of one-day culture of the microorganism at a concentration of  $5 \times 10^8$  CFU/ml on the partially de-epithelialized (approximately 1 cm<sup>2</sup>) cornea followed by coating for 24 hours with soft contact lens made of belafilcon A (water content: 36 %, oxygen permeability DK/t:110.0). In half of the cases, microbial biofilms were pre-grown on the contact lens surfaces via incubation into the broth culture of *P. aeruginosa* strain.

The treatment of keratitis was performed with ocular official forms of antibiotics: levofloxacin 0.5 % (5 mg/ml), ciprofloxacin 0.3 % (3 mg/ml), tobramycin 0.3 % (3 mg/ml). Their effectiveness against the strain of *P. aeruginosa* was previously proven *in vitro*. The animals were divided into three corresponding as for severity of keratitis groups, in each of which treatment was performed with one of the three above antibiotics. In half of the cases within each group, the antibiotic was combined with the ocular official form of dexamethoxine 0.02 % (0.2 mg/ml). The antibiotics were used in the mode of the most frequent instillation for the first 2 days after the onset of purulent keratitis, then 5 and 4 times a day.

The assessment included evaluation of clinical signs, culturing, ophthalmological examination of the cornea with fluorescein test and photofixation. Animals were removed from the study on days 7<sup>th</sup>, 10<sup>th</sup>, 14<sup>th</sup> (depending on the term of epithelialization of the affected cornea).

**Results.** Totally, 36 cases of purulent keratitis caused by *P. aeruginosa* in rabbits were modelled: 8 moderate (22.2 %), 13 semi-severe (36.6 %), 15 – severe (41.7 %).

Regardless of the chosen antibiotic, 10–12 hours after removal of the infected contact lens and the start of treatment, there was mostly a burst enhancement of the inflammatory process. Subsequently, the inflammatory signs showed a gradual attenuation with complete corneal epithelialization for 7<sup>th</sup>–8<sup>th</sup> days (for moderate keratitis), 10<sup>th</sup>–12<sup>th</sup> days (for severe and semi-severe in Group I and Group III) and (12<sup>th</sup>–14<sup>th</sup> days for severe and semi-severe in Group II). In comparison with the Group I (levofloxacin 0.5 %), the reducing of the corneal inflammation and lesion epithelialization within the Group II (ciprofloxacin 0.3 %), was 1–2 days delayed. The use of 0.3 % tobramycin (Group III) provided the highest control of purulent inflammation during the first three to four days, but further there was a slowdown in the positive dynamics and delayed epithelialization compared with Group I. For all study groups, the use of combination antibiotic therapy with dexamethoxine was accompanied by acceleration in regression of purulent-inflammatory lesions, especially due to a decrease in conjunctival response and pyorrhoea.

Among the outcomes, in most of cases (69.5 %) there were areas of corneal opaque turbidity of various sizes. Total leucoma was noted in more than a third of cases; always due to severe keratitis. It was managed to avoid the complications that could lead to eye loss (corneal abscess, perforation, malacia) in all animals. The culturing from the surface of the eye has shown a decrease from 100 % to 63.9 % in presence *P. aeruginosa* after 48 hours and the whole pathogen disappearance after 3 days of treatment. However, in 11 of 15 severe keratitis (30.6 % of total observations) microbiological tests of the cornea, obtained after removal of animals, were positive as for *P. aeruginosa* even after complete epithelialization.

**Conclusions.** Treatment of experimental keratitis caused by *P. aeruginosa*, with antibiotics, to which the sensitivity of pathogen is proven, in the mode of the earliest and most frequent instillations on the first 48 hours in combination with removal of purulent masses provides reducing of inflammation, restoring of cornea surface and prevents the most serious complications, e. g. corneal perforations and keratomalacia.

Early and active antimicrobial treatment with effective drugs led to the elimination of *P. aeruginosa* from the ocular surface in the first 2–3 days. However, in 30.6 % of cases there was no complete eradication of the pathogen from the cornea.

The choice of antibiotic from those, to which the sensitivity of *P. aeruginosa* is proven, did not have a significant influence on treatment outcome. Concomitant use of decamethoxine makes better effect and somewhat improves the results

**Keywords:** experimental keratitis, *Pseudomonas aeruginosa*, treatment, levofloxacin, ciprofloxacin, tobramycin, decamethoxine

## References

- Güell, J. L. (Ed.) (2015). Cornea. Vol. 6. ESASO Course Series. Basel: Karger, 127. doi: <http://doi.org/10.1159/000381489>
- Raphalyuk, S. Y. (2015). The possibility of metabolic correction of pathological disorders in induced keratitis in animals with dry eye syndrome. *Ophthalmology*, 2 (2), 211–220.
- Zamani, M., Panahi-Bazaz, M., Assadi, M. (2015). Corneal collagen cross-linking for treatment of non-healing corneal ulcers. *Journal of Ophthalmic and Vision Research*, 10 (1), 16–20. doi: <http://doi.org/10.4103/2008-322x.156087>
- Gokhale, N. (2008). Medical management approach to infectious keratitis. *Indian Journal of Ophthalmology*, 56 (3), 215–220. doi: <http://doi.org/10.4103/0301-4738.40360>
- Farias, R., Pinho, L., Santos, R. (2017). Epidemiological profile of infectious keratitis. *Revista Brasileira de Oftalmologia*, 76 (3), 116–120. doi: <http://doi.org/10.5935/0034-7280.20170024>
- Wang, M., Smith, W. A., Duncan, J. K., Miller, J. M. (2017). Treatment of *Pseudomonas* Keratitis by Continuous Infusion of Topical Antibiotics With the Morgan Lens. *Cornea*, 36 (5), 617–620. doi: <http://doi.org/10.1097/ico.0000000000001128>
- Tacconelli, E., Carrara, E., Savoldi, A., Kattula, D., Burkert, F. (2017). Global priority list of antibiotic-resistant bacteriatoguidere search, discovery, and development of new antibiotics World Health Organization. Available at: [https://www.who.int/medicines/publications/WHO-PPL-Short\\_Summary\\_25Feb-ET\\_NM\\_WHO.pdf](https://www.who.int/medicines/publications/WHO-PPL-Short_Summary_25Feb-ET_NM_WHO.pdf)
- Green, M., Apel, A., Stapleton, F. (2008). Risk Factors and Causative Organisms in Microbial Keratitis. *Cornea*, 27 (1), 22–27. doi: <http://doi.org/10.1097/ico.0b013e318156caf2>
- Jin, H., Parker, W. T., Law, N. W., Clarke, C. L., Gisseman, J. D., Pflugfelder, S. C. et. al. (2017). Evolving risk factors and antibiotic sensitivity patterns for microbial keratitis at a large county hospital. *British Journal of Ophthalmology*, 101 (11), 1483–1487. doi: <http://doi.org/10.1136/bjophthalmol-2016-310026>
- Zimmerman, A. B., Nixon, A. D., Rueff, E. M. (2016). Contact lens associated microbial keratitis: practical considerations for the optometrist. *Clinical Optometry*, 8, 1–12. doi: <http://doi.org/10.2147/opto.s66424>
- Dart, J. K. G., Seal, D. V. (1988). Pathogenesis and therapy of *pseudomonas aeruginosa* keratitis. *Eye*, 2 (S1), S46–S55. doi: <http://doi.org/10.1038/eye.1988.133>
- Rykov, S. A., Lemziakov, G. G., Bakbardina, L. M., Bakbardina, I. I. (2009). *Zabolievaniya rogovitsy*. Kyiv, 244.
- Lambert, R. J. W., Hanlon, G. W., Denyer, S. P. (2004). The synergistic effect of EDTA/antimicrobial combinations on *Pseudomonas aeruginosa*. *Journal of Applied Microbiology*, 96 (2), 244–253. doi: <http://doi.org/10.1046/j.1365-2672.2004.02135.x>
- Melezhyk, I. A., Yavorskaya, N. V., Shepelevich, V. V., Kokozay, V. N. (2013). The role of biofilms in *pseudomonas aeruginosa* endogenous infections. *Biulleten Orenburgskogo nauchnogo centra UrO RAN*, 3. Available at: [http://elmag.uran.ru:9673/magazine/Numbers/2013-3/Articles/5Melezhyk\(2013-3\).pdf](http://elmag.uran.ru:9673/magazine/Numbers/2013-3/Articles/5Melezhyk(2013-3).pdf)
- Mulcahy, L. R., Isabella, V. M., Lewis, K. (2013). *Pseudomonas aeruginosa* Biofilms in Disease. *Microbial Ecology*, 68 (1), 1–12. doi: <http://doi.org/10.1007/s00248-013-0297-x>
- O'Callaghan, R., Caballero, A., Tang, A., Bierdeman, M. (2019). *Pseudomonas aeruginosa* Keratitis: Protease IV and PASP as Corneal Virulence Mediators. *Microorganisms*, 7 (9), 281. doi: <http://doi.org/10.3390/microorganisms7090281>
- Marquart, M. E. (2011). Animal Models of Bacterial Keratitis. *Journal of Biomedicine and Biotechnology*, 2011, 1–12. doi: <http://doi.org/10.1155/2011/680642>
- McClellan, S., Jiang, X., Barrett, R., Hazlett, L. D. (2015). High-Mobility Group Box 1: A Novel Target for Treatment of *Pseudomonas aeruginosa* Keratitis. *The Journal of Immunology*, 194 (4), 1776–1787. doi: <http://doi.org/10.4049/jimmunol.1401684>
- Malachkova, N. V., Kryvetska, N. V., Vovk, I. M., Kryvetskiy, V. F. (2020). Pat. No. 141155. Method of modeling of pseudomonad keratitis in rabbits. MPK: C12Q 1/00, C12R 1/385. G09B 23/28. No. u201908915. declared: 23.07.19; published: 25.03.2020, No. 6.
- Malachkova, N. V., Kryvetska, N. V., Vovk, I. M., Kryvetskiy, V. F. (2020). Pat. No. 141156. Method of modeling of associated with contact lens pseudomonad keratitis in rabbits. MPK: C12Q 1/00, G09B 23/28, C12R 1/385. declared: 23.07.19; published: 25.03.2020, No. 6.
- Malachkova, N. V., Kryvetska, N. V., Vovk, I. M., Kovalenko, I. M., Kryzhanovskaya, A. V. (2020). Models of experimental pseudomonas keratitis: microbiological and clinical aspects. Reports of Vinnytsia National Medical University, 24 (1), 129–133. doi: [http://doi.org/10.31393/reports-vnmedical-2020-24\(1\)-25](http://doi.org/10.31393/reports-vnmedical-2020-24(1)-25)
- Vovk, I. M., Kryvetska, N. V., Burkot, V. M., Dudar, A. O., Kulik, A. V. (2020). Microbiological grounds for antimicrobial treatment of experimental pseudomonad keratitis. Reports of Vinnytsia National Medical University, 24 (1), 114–117. doi: [http://doi.org/10.31393/reports-vnmedical-2020-24\(1\)-21](http://doi.org/10.31393/reports-vnmedical-2020-24(1)-21)
- Lin, A., Rhee, M. K., Akpek, E. K., Amescua, G., Farid, M., Garcia-Ferrer, F. J. et. al. (2019). Bacterial Keratitis Preferred Practice Pattern®. *Ophthalmology*, 126 (1), 1–55. doi: <http://doi.org/10.1016/j.ophtha.2018.10.018>
- Rasamiravaka, T., Labtani, Q., Duez, P., El Jaziri, M. (2015). The Formation of Biofilms by *Pseudomonas aeruginosa*: A Review of the Natural and Synthetic Compounds Interfering with Control Mechanisms. *BioMed Research International*, 2015, 1–17. doi: <http://doi.org/10.1155/2015/759348>

25. Nazarchuk, O., Chereszniuk, I., Nazarchuk, G., Palii, D. et. al. (2019). Current antiseptics: a study on their antimicrobial activity and toxic effects on the corneal epithelium. *Oftalmologicheskii Zhurnal*, 80 (3), 26–31. doi: <http://doi.org/10.31288/oftalmolzh201932631>

26. Bezwada, P., Clark, L. A., Schneider, S. (2007). Intrinsic cytotoxic effects of fluoroquinolones on human corneal keratocytes and endothelial cells. *Current Medical Research and Opinion*, 24 (2), 419–424. doi: <http://doi.org/10.1185/030079908x261005>

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## STUDY OF THE LACTOSE INTOLERANCE GENETIC ASPECTS AMONG THE POPULATION AND PATIENTS WITH HIP FRACTURE IN EASTERN UKRAINE

p. 29-33

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*About 75 % of the world's population loses ability to lactose tolerance in adulthood. The study of tolerance to lactose (LT) or lactase persistence (LP) in Ukrainians is necessary to identify the association of this trait with multifactorial pathologies, the formation of risk groups and food culture development.*

*The aim of this study was to evaluate the genetic aspects of lactose deficiency in population and its relation with human pathologies on the example of a hip fracture.*

**Material and methods:** lactose intolerance phenotype was found in 9.7 % of Ukrainians and lactose persistence amounted to 69.4 %. In group of patients with fractures of the proximal femur frequencies were: 13910T – 0.34, 13910C – 0.66, 22018A – 0.35, 22018G – 0.65. Distribution of genotypes was 0 % : 67.6 % : 32.4 % for TT:CT:CC and 2.9 % : 64.7 % : 32.4 %, for AA:GA:GG. Deviation from the Hardy-Weinberg equilibrium was not observed.

**Results:** when analyzed genotypes for both SNPs, we found the ratio – CTA:CTGA:CCGA:CCGG being 2.9 % : 61.7 % : 2.9 % : 32.5 %. The linkage disequilibrium was estimated,  $D'$  ( $r^2$ ) for SNPs analyzed was 0.209 (0.554). A half of patients had osteoporosis or osteopenia ( $n=50.0$  %), but these pathologies were not observed in patients with the TTAA genotype. The param-

eters BMD of CTGA patients were  $4049.8 \pm 38.7$  and  $4109.6 \pm 30.0$  for CCGG patients.

**Conclusion:** all patients with osteoporosis had a genotype CTGA, average value of BMD was  $3795.6 \pm 43.8$ . CTGA-patients had following gastrointestinal tract disorders: gastritis (33.3 %), peptic ulcer (9.5 %), duodenal ulcer (9.5 %), stomach cancer (9.5 %). Heterozygous genotypes significantly increase five times the risk of gastrointestinal pathology in patients with fractures of the proximal femur

**Keywords:** lactase persistence, lactose intolerance, hip fracture, osteoporosis, genotypes, bone mineral density

## References

1. Kettunen, J., Silander, K., Saarela, O., Amin, N., Muller, M., Timpson, N. et. al. (2009). European lactase persistence genotype shows evidence of association with increase in body mass index. *Human Molecular Genetics*, 19 (6), 1129–1136. doi: <http://doi.org/10.1093/hmg/ddp561>
2. Bonjour, J.-P. (2013). Nutritional disturbance in acid–base balance and osteoporosis: a hypothesis that disregards the essential homeostatic role of the kidney. *British Journal of Nutrition*, 110 (7), 1168–1177. doi: <http://doi.org/10.1017/s0007114513000962>
3. Enattah, N. S., Sahi, T., Savilähti, E., Terwilliger, J. D., Peltonen, L., Järvelä, I. (2002). Identification of a variant associated with adult-type hypolactasia. *Nature Genetics*, 30 (2), 233–237. doi: <http://doi.org/10.1038/ng826>
4. Ji, J., Sundquist, J., Sundquist, K. (2014). Lactose intolerance and risk of lung, breast and ovarian cancers: aetiological clues from a population-based study in Sweden. *British Journal of Cancer*, 112 (1), 149–152. doi: <http://doi.org/10.1038/bjc.2014.544>
5. Somekawa, Y., Chiguchi, M., Ishibashi, T., Aso, T. (2001). Soy intake related to menopausal symptoms, serum lipids, and bone mineral density in postmenopausal Japanese women. *Obstetrics & Gynecology*, 97 (1), 109–115. doi: [http://doi.org/10.1016/s0029-7844\(00\)01080-2](http://doi.org/10.1016/s0029-7844(00)01080-2)
6. Wagh, K., Bhatia, A., Alexe, G., Reddy, A., Ravikumar, V., Seiler, M. et. al. (2012). Lactase Persistence and Lipid Pathway Selection in the Maasai. *PLoS ONE*, 7 (9), e44751. doi: <http://doi.org/10.1371/journal.pone.0044751>
7. Albuquerque, D., Nóbrega, C., Manco, L. (2013). The lactase persistence -13910C>T polymorphism shows indication of association with abdominal obesity among Portuguese children. *Acta Paediatrica*, 102 (4), e153–e157. doi: <http://doi.org/10.1111/apa.12134>
8. Guo, J., Astrup, A., Lovegrove, J. A., Gijsbers, L., Givens, D. I., Soedamah-Muthu, S. S. (2017). Milk and dairy consumption and risk of cardiovascular diseases and all-cause mortality: dose–response meta-analysis of prospective cohort studies. *European Journal of Epidemiology*, 32 (4), 269–287. doi: <http://doi.org/10.1007/s10654-017-0243-1>
9. Fenton, T. R., Tough, S. C., Lyon, A. W., Eliasziw, M., Hanley, D. A. (2011). Causal assessment of dietary acid load and bone disease: a systematic review & meta-analysis applying Hill's epidemiologic criteria for causality. *Nutrition Journal*, 10(1). doi: <http://doi.org/10.1186/1475-2891-10-41>
10. Shams-White, M. M., Chung, M., Fu, Z., Insogna, K. L., Karlsen, M. C., LeBoff, M. S. et. al. (2018). Animal versus plant protein and adult bone health: A systematic review and meta-analysis from the National Osteoporosis Foundation. *PLOS ONE*, 13 (2), e0192459. doi: <http://doi.org/10.1371/journal.pone.0192459>
11. Kalchenko, A., Babalyan, V., Hurbanova, T., Maznyakov, S. (2016). Surgical treatment of proximal femur osteoporotic fractures (literature review). *Orthopaedics, Traumatology and Prosthetics*, 2, 111–119. doi: <http://doi.org/10.15674/0030-598720162111-119>

12. Rocha-Braz, M. G. M., Ferraz-de-Souza, B. (2016). Genetics of osteoporosis: searching for candidate genes for bone fragility. *Archives of Endocrinology and Metabolism*, 60 (4), 391–401. doi: <http://doi.org/10.1590/2359-3997000000178>

13. Kuchay, R. A. H., Anwar, M., Thapa, B. R., Mahmood, A., Mahmood, S. (2012). Correlation of G/A -22018 single-nucleotide polymorphism with lactase activity and its usefulness in improving the diagnosis of adult-type hypolactasia among North Indian children. *Genes & Nutrition*, 8 (1), 145–151. doi: <http://doi.org/10.1007/s12263-012-0305-7>

14. Coelho, M., Luiselli, D., Bertorelle, G., Lopes, A. I., Seixas, S., Destro-Bisul, G., Rocha, J. (2005). Microsatellite variation and evolution of human lactase persistence. *Human Genetics*, 117 (4), 329–339. doi: <http://doi.org/10.1007/s00439-005-1322-z>

15. Hodges, J., Cao, S., Cladis, D., Weaver, C. (2019). Lactose Intolerance and Bone Health: The Challenge of Ensuring Adequate Calcium Intake. *Nutrients*, 11 (4), 718. doi: <http://doi.org/10.3390/nu11040718>

16. Fedota, O. M., Babalian, V. O., Boroznets, V. V., Mazniakov, S. M., Puzik, N. G. (2019). Lactose intolerance and its association with the exogamy degree among the population of eastern Ukraine. *Faktori Eksperimental'noi Evolucii Organizmiv*, 24, 249–253. doi: <http://doi.org/10.7124/feeo.v24.1110>

17. Iacone, R., Scanzano, C., D'Isanto, A., Vitalone, A., Frangipane, I., D'Angeli, M. et. al. (2018). Prediction of Renal Acid Load in Adult Patients on Parenteral Nutrition. *Pharmaceutics*, 10 (2). doi: <http://doi.org/10.3390/pharmaceutics10020043>

18. Baranov, V. S. (2009). Geneticpassport-the basis of individual and predictive medicine. Saint-Petersburg, 528.

19. Fedota, O. M., Babalian, V. O., Mitiohlo, L. V., Mazniakov, S. M., Valilshchikov, M. V., Tyzhnenko, T. V., Ruban S. Yu. (2017). Bone mineral density in evaluation of the productive traits and reproductive health of dairy cows. *Journal for veterinary medicine, biotechnology and biosafety*, 3 (4), 16–22. Available at: [http://nbuv.gov.ua/UJRN/jvmbb\\_2017\\_3\\_4\\_6](http://nbuv.gov.ua/UJRN/jvmbb_2017_3_4_6)

20. Larsson, S. C., Orsini, N., Wolk, A. (2006). Milk, milk products and lactose intake and ovarian cancer risk: A meta-analysis of epidemiological studies. *International Journal of Cancer*, 118 (2), 431–441. doi: <http://doi.org/10.1002/ijc.21305>

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## RADIONUCLIDE DIAGNOSTICS OF JOINT INJURY IN PATIENTS WITH DIABETES

p. 34-40

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*There are very few convincing effective ways of early diagnosis of joint lesions today. Radiological changes in the early stages of the disease are absent, other laboratory tests are not specific and do not always correlate with the activity of the inflammatory process.*

*The aim of our work was to study with the help of radionuclide method the state of arterial and venous blood flow, the presence of inflammatory processes in the knee and ankle joints in patients with type 1 and 2 diabetes mellitus.*

*Materials and methods.* 152 patients with diabetes were examined. Depending on the type of diabetes, patients were divided into two groups – 58 patients were with type 1 diabetes (19 men and 39 women), with type 2 diabetes – 94 patients (19 men and 51 women). The mean age of patients in the type 1 diabetes mellitus was 42.3±1.8 years, the duration of diabetes mellitus was 21.1±1.3 years. In the group of type 2 diabetes mellitus, the average age of patients was 61.3±1.0 years, the duration of diabetes mellitus was 13.1±0.9 years. Among the examined patients with type 1 diabetes mellitus, arthropathy was diagnosed in 34 (58.6 %), the control group was patients without joint damage – 24 (41.4 %). In patients with type 2 diabetes mellitus arthropathy was found in 68 (72.3 %) subjects, joint pathology was not in 26 (27.7 %) patients. The method of radionuclide diagnostics of the lower extremities consisted of two methods: radionuclide angiography and radionuclide scintigraphy.

*Results.* At carrying out radionuclide researches of hemodynamics at patients with a diabetes mellitus 1 type with arthropathies in comparison with patients without defeat of joints there are no changes of a blood-groove in vessels of large and average calibre and also arterioles and capillaries of lower extremities in the presence of the expressed venous outflow disturbances. For patients with type 2 diabetes mellitus with arthropathies, a significant slowing of blood flow velocity in large and medium-sized vessels, as well as arterioles and capillaries was found, which is accompanied by impaired venous outflow.

*Conclusions.* The applied method of radionuclide diagnostics to assess the state of blood flow and the presence of diabetes-associated arthropathy of the knee and ankle joints with a single injection of radiopharmaceutical is one of the most informative methods of early diagnosis of this pathology, and disorders of venous outflow may be a differential criterion for diabetes-associated arthropathy in patients with type 1 and type 2 diabetes

**Keywords:** diabetes mellitus, diabetes-associated arthropathy, radiosintigraphy, vascular disorders, risk factors, hemodynamics

## References

1. Vartanian, K. F. (2003). Kliniko-dagnosticheskie aspekty osteopatii pri sakharnom diabete. *Rossiiskie medicinskie vesti*, 3, 39–46.

2. Voloshyna, L. O., Kovalyshyn, I. R., Marchuk, Yu. F. (2013). Osteoartroz i tsukrovyyi diabet: epidemiolohichni osoblyvosti, kliniko-patohenetychni, henderni ta vikovi aspekty. *Mizhnarodnyi endokrynolohichniy zhurnal*, 6, 93–94.

3. Wang, Y., Dawson, C., Hanna, F., Fairley, J., Cicuttini, F. M. (2015). Association between popliteal artery wall thickness and knee cartilage volume loss in community-based middle-aged women without clinical knee disease. *Maturitas*, 82 (2), 222–227. doi: <http://doi.org/10.1016/j.maturitas.2015.07.010>

4. Slavnov, V. M., Savvitskyi, S. Yu. (2011). Radionuklidni metody v otsinitsi efektyvnosti medykamentoznoho likuvannia diabetichnykh osteoartropatii i anhiopatii nyzhnikh kintsivok. *Ukrainskyi radiolohichnyi zhurnal*, 3, 320–321.

5. Rehling, T., Björkman, A.-S. D., Andersen, M. B., Ekholm, O., Molsted, S. (2019). Diabetes Is Associated with Musculoskeletal Pain, Osteoarthritis, Osteoporosis, and Rheumatoid Arthritis. *Journal of Diabetes Research*, 2019, 1–6. doi: <http://doi.org/10.1155/2019/6324348>

6. Findlay, D. M. (2007). Vascular pathology and osteoarthritis. *Rheumatology*, 46 (12), 1763–1768. doi: <http://doi.org/10.1093/rheumatology/kem191>

7. Burgener, F. A., Kormano, M., Pudas, T. (2014). Luchevaia diagnostika zabolevanii kostei i sustavov: rukovodstvo, atlas: bolee 1000 rentgenogramm. Moscow: GEOTAR-Media, 539.

8. Ozgul, A., Yasar, E., Arslan, N., Balaban, B., Taskayntan, M. A., Tezel, K. et. al. (2008). The comparison of ultrasonographic and scintigraphic findings of early arthritis in revealing rheumatoid arthritis according to criteria of American College of Rheumatology. *Rheumatology International*, 29 (7), 765–768. doi: <http://doi.org/10.1007/s00296-008-0765-7>

9. Bergaliev, A. N., Fadeev, N. P., Pozdeev, A. P. (2011). Rol polifaznoi osteoscintigrafii v ocenke sostoianiiia perfuzionno-metabolicheskikh processov pri zabolevaniiah oporno-dvigatel'nogo apparata u detei. *Genii ortopedii*, 2, 140–147.

10. Kim, J. Y., Choi, Y. Y., Kim, C. W., Sung, Y.-K., Yoo, D.-H. (2016). Bone Scintigraphy in the Diagnosis of Rheumatoid Arthritis: Is There Additional Value of Bone Scintigraphy with Blood Pool Phase over Conventional Bone Scintigraphy? *Journal of Korean Medical Science*, 31 (4), 502–509. doi: <http://doi.org/10.3346/jkms.2016.31.4.502>

11. Zhuravlova, L. V., Oliinyk, M. O. (2015). Osteoartroz ta tsukrovyyi diabet 2 typu: spilni lanky patohenezu. *Endokrynolohiia*, 20 (1), 447–451.

12. Zubareva, E. V., Lesniak, O. M. (2016). Vliiaet li varikozaia bolezn nizhnikh konechnosti na proiavlennia osteoartroza kolennykh sustavov? *Lechaschii vrach*, 12, 64–67.

13. Lesniak, O. M., Zubareva, E. V., Goncharova, M. G., Maksimov, D. M. (2017). Patologiia ven nizhnikh konechnosti pri pervichnom osteoartroze kolennykh sustavov. *Terapevticheskii arkhiv*, 89 (5), 53–59.

14. Conaghan, P. G., Vanharanta, H., Dieppe, P. A. (2005). Is progressive osteoarthritis an atheromatous vascular disease? *Annals of the Rheumatic Diseases*, 64 (11), 1539–1541. doi: <http://doi.org/10.1136/ard.2005.039263>

15. Masaoka, S. (2001). Evaluation of arterial obstructive leg and foot disease by three-phase bone scintigraphy. *Annals of Nuclear Medicine*, 15 (3), 281–287. doi: <http://doi.org/10.1007/bf02987847>

16. Scheglov, E. A., Vezikova, N. N. (2012). Osobennosti klinicheskoi kartiny u pacientov osteoartrozom kolennykh sustavov i sochetannym porazheniem ven nizhnikh konechnosti. *Sovremennye problemy nauki i obrazovaniia*, 1, 65–71.

17. Liubarskii, M. S., Mustafaev, N. R. (2011). Izmeneniia gemolimfocirkuliacii nizhnikh konechnosti pod vliianiem fizioterapevticheskikh metodov v lechenii osteoartroza kolennykh. *Sovremennye problemy nauki i obrazovaniia*, 1, 23–26.

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## MICROHARDNESS OF FIBERGLASS – REINFORCED PHOTOCOMPOSITE MATERIAL UNDER DIFFERENT CONDITIONS OF LIGHT POLIMERIZATION

p. 40-44

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*The aim:* to study in laboratory conditions the microhardness of a glass-fiber-reinforced photocomposite under different modes of light exposure at different times.

*Materials and methods.* The microhardness of the everX Posterior, GC glass-fiber-reinforced photocomposite was investigated on 60 samples using a PMT-3 microhardness meter on three sample surfaces within 1 hour, 1 day and 7 days after polymerization. Samples of a cylindrical shape with a height of 3 mm of group 1 were irradiated with the light flux of a photopolymerizer in the “soft start” mode, samples of group 2 were polymerized with light of constant high intensity 1400 mW/cm<sup>2</sup>.

*Research results.* After 1 hour, the microhardness on the surface closest to the LED was 87.34±1.21 kgf/mm<sup>2</sup> in samples of group 1, 102.0±0.94 kgf/mm<sup>2</sup> (p<0.05) in group 2, and 70.98±1.23 kgf/mm<sup>2</sup> (the lowest indicator) and 90.65±1.12 kgf/mm<sup>2</sup> (p<0.05). After 1 day, on the nearest surface, the microhardness increased in the samples of group 1 to 97.03±1.25 kgf/mm<sup>2</sup>, group 2 – to 114.61±1.13 kgf/mm<sup>2</sup> (p<0.05), on the most distant – up to 75.95±1.11 kgf/mm<sup>2</sup> and 99.83±1.24 kgf/mm<sup>2</sup> (p<0.05), respectively. At 7 days, the indicators on the first surface in group 1 were 104.64±1.23 kgf/mm<sup>2</sup>, in 2 – 123.35±1.15 kgf/mm<sup>2</sup> (p<0.05), on the other surface – 80.25±1.48 kgf/mm<sup>2</sup> and 107.53±0.92 kgf/mm<sup>2</sup> (p<0.05). The growth of microhardness on these surfaces for the entire period was 16.5 % and 11.6 % in the samples of group 1, and 17.3 % and 15.7 % in group 2.

*Conclusions.* The light flux of constant high intensity provides statistically significantly (p<0.05) higher microhardness indices of the glass-fiber reinforced photocomposite on all surfaces of the samples than the light exposure in the “soft start” mode. In the direct restoration of teeth, it is necessary to reduce the thickness of the photocomposite layer for “soft start” polymerization

**Keywords:** glass-fiber reinforced photocomposite, microhardness, polymerization, luminous flux, intensity, “soft start”

### References

1. Borisenko, A. V., Nespryad'ko, V. P., Borisenko, D. A. (2015). Kompozitsionnye plombirovochnye i oblitsovochnye materialy. Kyiv: VSI «Meditsina», 320.

2. Adalaev, H. I. (2017). Zhidkotekuchie kompozitsionnye materialy svetovogo otverzheniya. *Bulletin of Medical Internet Conferences*, 7 (10), 1554–1555.

3. Gryttsner, A. (2011). Tekuchiy kompozit SDR – umniy zamenitel' dentina. *DentArt*, 2, 45–52.



4. Wolff, D., Geiger, S., Ding, P., Staehle, H. J., Frese, C. (2012). Analysis of the interdiffusion of resin monomers into pre-polymerized fiber-reinforced composites. *Dental Materials*, 28 (5), 541–547. doi: <https://doi.org/10.1016/j.dental.2011.12.001>

5. Miletich, I. (2018). Sovremennyye resheniya dlya pryamyh restavratsiy zubov distal'noy gruppy. *Glavniy vrach Yuga Rossii*, 61, 6–9.

6. Garoushi, S., Vallittu, P. K., Watts, D. C., Lassila, L. V. J. (2008). Effect of nanofiller fractions and temperature on polymerization shrinkage on glass fiber reinforced filling material. *Dental Materials*, 24 (5), 606–610. doi: <https://doi.org/10.1016/j.dental.2007.06.020>

7. EverX Posterior. Kompozyt dlia zamishchennia dentyu, pidsylenyi voloknom. Available at: <https://kristar.ua/upload/iblock/c18/c185a519bff1c413136026feb1f1913d.pdf>

8. Udod, O. A., Bekuzarova, K. I. (2018). Study of nanophotocomposite material microhardness under various hardening conditions. *Bulletin of Problems Biology and Medicine*, 4.3 (141), 260. doi: <https://doi.org/10.29254/2077-4214-2017-4-3-141-260-263>

9. Udod, O. A., Roman, O. B. (2020). Doslidzhennia hlybiny polimeryzatsiyi fotokompozytsiynykh materialiv. *Mater. mizhnar. nauk.-prakt. konf. «Medychna nauka ta praktyka na suchasnomu istorychnomu etapi»*. Kyiv, 116–118.

10. GOST 9450-76. Izmerenie mikrotverdosti vдавlivanem almaznyh nakonechnikov (1993). Moscow: Izd-vo standartov, 35.

11. Udod, O. A., Bakuzarova, H. I. (2018). Investigating the Intensity Dynamics of the Photopolymerizer Light Flux in Restorative Materials. *Ukrainskyi zhurnal medytsyny, biolohiyi ta sportu*, 3 (2), 171–174. doi: <https://doi.org/10.26693/jmbs03.02.171>

12. Maniuh, H. Yu., Maksymiv, O. O., Rozhko, V. I. (2012). A modern view of photocomposite filling materials and their features in case of restoring defects of the crown part of the teeth. *Bukovynskyi medychnyi visnyk*, 16 (1 (61)), 166–170.

13. Garoushi, S., Tanner, J., Vallittu, P., Lassila, L. (2012). Preliminary Clinical Evaluation of Short Fiber-Reinforced Composite Resin in Posterior Teeth: 12-Months Report. *The Open Dentistry Journal*, 6 (1), 41–45. doi: <https://doi.org/10.2174/1874210601206010041>

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#### ANALYSIS OF BLOOD GAS COMPOSITION INDICATORS IN PREMATURE BABIES WITH NEONATAL SEPSIS

p. 45-48

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*The aim of the study was analysis of oxygen status of patients with neonatal sepsis during monitoring of premature infants.*

**Materials and methods.** Indicators of oxygen status of capillary blood were assessed in premature infants with neonatal sepsis who were treated at the Regional Perinatal Center in Khmelnytsky during 2017–2018. The study included mostly deeply premature babies, including those with extremely low weight. A full range of clinical and laboratory studies, including determination of oxygen status of capillary blood.

**Research results.** Our results allow us to conclude that while such indicators as pH, pO<sub>2</sub>, BEb3, % SO<sub>2</sub> do not change significantly, the trouble can be detected using indicators such as AdDO<sub>2</sub>, RI, PI, which reflect the state of oxygenation with impaired ventilation. -perfusion ratios, increased blood shunting and the processes of delivery and consumption of oxygen at the tissue level. The calculated indicator “PIP×FiO<sub>2</sub>” was less sensitive to assess changes in the lungs and did not correspond to the clinical picture of the severity of the patient's condition.

**Conclusion.** Thus, the determination of AdDO<sub>2</sub>, RI, PI capillary blood can be used as informative indicators for determining the oxygen status in premature infants, which is of significant practical importance in terms of limited monitoring of the gas composition of arterial blood

**Keywords:** neonatal sepsis, premature infants, oxygen status indicators, respiratory index

#### References

1. Parshin, Y. V., Aleksandrovich, Y. S., Kushnerik, L. A., Blinov, S. A., Pshenisnov, K. V., Nurmagambetova, B. K. (2010). Oxygen Status Parameters as Markers of Renal Dysfunction in Neonatal Infants with Critical Status. *General Reanimatology*, 6 (2), 62–67. doi: <http://doi.org/10.15360/1813-9779-2010-2-62>

2. Arayici, S., Şimşek, G. K., Canpolat, F. E., Oncel, M. Y., Uras, N., Oğuz, S. S. (2019). Can Base Excess be Used for Prediction to Early Diagnosis of Neonatal Sepsis in Preterm Newborns? *Mediterranean journal of hematology and infectious diseases*, 11 (1), e2019014. doi: <http://doi.org/10.4084/mjihid.2019.014>

3. Kassie, D. G., Tewolde, A. W. S., Bogale, W. A. (2020). Premature Rupture of Membrane and Birth Asphyxia Increased Risk of Neonatal Sepsis Among Neonates in the Neonatal Intensive Care Unit at the University of Gondar Specialized Referral Hospital, Northwest Ethiopia. *The Pediatric Infectious Disease Journal*, 5 (1), 1. doi: <http://doi.org/10.36648/2573-0282.5.1.68>

4. Perestoronina, M. V. (2015). Comparison of oxygen indicators in capillary blood full term newborns and infants with extremely low birth weight. *Omskii nauchnii vestnik*, 2 (144), 113–115.

5. Shkurupii, D. A. (2013). Zahalni tendentsii klinichnoho perebihu syndromu poliorrhanoi nedostatnosti u novonarodzhennykh. *Bil, Znebolivannia i intensyvna terapiia*, 1, 46–51.

6. Petrenko, Iu. V., Ivanov, D. O., Kurzina, E. A. (2011). Ocenka organoz nedostatochnosti u novorozhdennykh. *Biulleten federalnogo centra serdca, krovi i endokrinologii im. V. A. Almazova*, 43–50.

7. Perestoronina, M. V., Korpacheva, O. V., Palyanov, S. V., Dolgikh, V. T. (2015). The parameters of the oxygen status in the assessment of prognosis of a hemodynamically significant patent ductus arteriosus preterm neonatal infants. *General Reanimatology*, 11 (2), 35–41. doi: <http://doi.org/10.15360/1813-9779-2015-2-35-41>

8. Tan, R. N. G. B., Pauws, S. C., van Loon, E., Smits, V. E. H. J., Lopriore, E., te Pas, A. B. (2018). Correlation and Interchangeability of Venous and Capillary Blood Gases in Non-Critically Ill Neonates. *Frontiers in Pediatrics*, 6. doi: <http://doi.org/10.3389/fped.2018.00089>

9. Andersen, C. C., Hodyl, N. A., Kirpalani, H. M., Stark, M. J. (2017). A Theoretical and Practical Approach to De-

fining “Adequate Oxygenation” in the Preterm Newborn. *Pediatrics*, 139 (4), e20161117. doi: <http://doi.org/10.1542/peds.2016-1117>

10. Kayton, A., Timoney, P., Vargo, L., Perez, J. A. (2018). A Review of Oxygen Physiology and Appropriate Management of Oxygen Levels in Premature Neonates. *Advances in Neonatal Care*, 18 (2), 98–104. doi: <http://doi.org/10.1097/anc.0000000000000434>

11. Grizelj, R., Bojanić, K., Pritišanac, E., Luetić, T., Vučković, J., Weingarten, T. N. et. al. (2016). Survival prediction of high-risk outborn neonates with congenital diaphragmatic hernia from capillary blood gases. *BMC Pediatrics*, 16 (1). doi: <http://doi.org/10.1186/s12887-016-0658-y>

12. Singer, M., Deutschman, C. S., Seymour, C. W., Shankar-Hari, M., Annane, D., Bauer, M. et. al. (2016). The Third International Consensus Definitions for Sepsis and Septic Shock (Sepsis-3). *JAMA*, 315 (8), 801–810. doi: <http://doi.org/10.1001/jama.2016.0287>

13. Ruan, L., Chen, G.-Y., Liu, Z., Zhao, Y., Xu, G.-Y., Li, S.-F. et. al. (2018). The combination of procalcitonin and C-reactive protein or presepsin alone improves the accuracy of diagnosis of neonatal sepsis: a meta-analysis and systematic review. *Critical Care*, 22 (1). doi: <http://doi.org/10.1186/s13054-018-2236-1>

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#### MODERN APPROACHES TO DIAGNOSIS AND TREATMENT OF EARLY-ONSET NEONATAL SEPSIS

p. 49-52

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*Early-onset neonatal sepsis (EOS) remains the leading cause of morbidity and mortality, especially among premature babies. Therefore, accurate diagnosis, prompt monitoring of the course of the disease and an effective therapeutic strategy are a guarantee of improving the quality of medical care for newborns, as well as an important reserve for reducing perinatal losses.*

*The aim of the work* to summarize modern views on the diagnosis and treatment of early-onset neonatal sepsis in newborns with perinatal pathology.

*Results and their discussion.* An analytical review of modern literature data on problematic issues of EOS diagnostics is carried out. The approaches are stated and the main directions of EOS management at the present stage are formulated. It was revealed that the lack of a consensus definition of neonatal sepsis, the nonspecificity of clinical data is a limiting factor in the timely accurate identification of the septic process. Insufficient

*diagnostic value of existing laboratory tests leads to defects in the registry and monitoring of this disease, the inability to conduct an objective assessment of the existing epidemiological and microbiological situation in the health care system and the subsequent effective implementation of preventive measures.*

**Conclusions.** In the conditions of modern scientific realities, none of the existing laboratory tests can sufficiently reliably confirm or deny the presence of EOS in a newborn child, which determines the search for new promising laboratory tests with high diagnostic and prognostic potential. The optimal treatment strategy for newborns with EOS is broad-spectrum antibiotics

**Keywords:** newborns, neonatal sepsis, early-onset neonatal sepsis, diagnosis, antibacterial therapy

#### References

1. Benitz, W. E., Achten, N. B. (2020). Finding a role for the neonatal early-onset sepsis risk calculator. *EClinicalMedicine*, 19, 100255. doi: <http://doi.org/10.1016/j.eclinm.2019.100255>

2. Puopolo, K. M. (2019). Neonatal sepsis evaluation across the pond. *Archives of Disease in Childhood – Fetal and Neonatal Edition*, 105 (2), 116–117. doi: <http://doi.org/10.1136/archdischild-2019-317840>

3. Puopolo, K. M., Benitz, W. E., Zaoutis, T. E. (2018). Management of Neonates Born at  $\leq 34$  6/7 Weeks' Gestation With Suspected or Proven Early-Onset Bacterial Sepsis. *Pediatrics*, 142 (6), e20182896. doi: <http://doi.org/10.1542/peds.2018-2896>

4. Watal, C., Kler, N., Oberoi, J. K., Fursule, A., Kumar, A., Thakur, A. (2019). Neonatal Sepsis: Mortality and Morbidity in Neonatal Sepsis due to Multidrug-Resistant (MDR) Organisms: Part 1. *The Indian Journal of Pediatrics*, 87 (2), 117–121. doi: <http://doi.org/10.1007/s12098-019-03106-z>

5. Giannoni, E., Agyeman, P. K. A., Stocker, M., Posfay-Barbe, K. M., Heininger, U., Spycher, B. D. et. al. (2018). Neonatal Sepsis of Early Onset, and Hospital-Acquired and Community-Acquired Late Onset: A Prospective Population-Based Cohort Study. *The Journal of Pediatrics*, 201, 106–114.e4. doi: <http://doi.org/10.1016/j.jpeds.2018.05.048>

6. Schrag, S. J., Farley, M. M., Petit, S., Reingold, A., Weston, E. J., Pondo, T. et. al. (2016). Epidemiology of Invasive Early-Onset Neonatal Sepsis, 2005 to 2014. *Pediatrics*, 138 (6), e20162013. doi: <http://doi.org/10.1542/peds.2016-2013>

7. Heorhiants, M. A., Zhovnir, V. A., Korsunov, V. A., Dmytriiev, D. V., Posternak, H. I., Snisar, V. I., Shchurovska, I. P. (2017). Unified clinical protocol of emergency, primary, secondary (specialized) and tertiary (highly specialized) medical care and intensive care Septic shock in children (project) (part 2). *Pain, Anesthesia and Intensive Care*, 4 (81), 11–26. doi: [http://doi.org/10.25284/2519-2078.4\(81\).2017.119236](http://doi.org/10.25284/2519-2078.4(81).2017.119236)

8. Wagstaff, J. S., Durrant, R. J., Newman, M. G., Eason, R., Ward, R. M., Sherwin, C. M. T., Enioutina, E. Y. (2019). Antibiotic Treatment of Suspected and Confirmed Neonatal Sepsis Within 28 Days of Birth: A Retrospective Analysis. *Frontiers in Pharmacology*, 10. doi: <http://doi.org/10.3389/fphar.2019.01191>

9. Zea-Vera, A., Ochoa, T. J. (2015). Challenges in the diagnosis and management of neonatal sepsis. *Journal of Tropical Pediatrics*, 61 (1), 1–13. doi: <http://doi.org/10.1093/tropej/fmu079>

10. Nikonov, V. V., Sokolov, A. S., Feskov, A. E. (2017). Sepsis from antiquity to contemporaneity. A view through the centuries. *Emergency Medicine*, 3 (82), 73–81. doi: <http://doi.org/10.22141/2224-0586.3.82.2017.102327>

11. Reinhart, K., Bauer, M., Riedemann, N. C., Hartog, C. S. (2012). *New Approaches to Sepsis: Molecular Diagnostics and*

Biomarkers. *Clinical Microbiology Reviews*, 25 (4), 609–634. doi: <http://doi.org/10.1128/cmr.00016-12>

12. Singer, M., Deutschman, C. S., Seymour, C. W., Shankar-Hari, M., Annane, D., Bauer, M. et al. (2016). The Third International Consensus Definitions for Sepsis and Septic Shock (Sepsis-3). *JAMA*, 315 (8), 801–810. doi: <http://doi.org/10.1001/jama.2016.0287>

13. Wynn, J. L. (2016). Defining neonatal sepsis. *Current Opinion in Pediatrics*, 28 (2), 135–140. doi: <http://doi.org/10.1097/mop.0000000000000315>

14. Polin, R. A. (2012). Management of Neonates With Suspected or Proven Early-Onset Bacterial Sepsis. *Pediatrics*, 129 (5), 1006–1015. doi: <http://doi.org/10.1542/peds.2012-0541>

15. Johansson Gudjónsdóttir, M., Elfvin, A., Hentz, E., Adlerberth, I., Tessin, I., Trollfors, B. (2019). Changes in incidence and etiology of early-onset neonatal infections 1997–2017 – a retrospective cohort study in western Sweden. *BMC Pediatrics*, 19 (1). doi: <http://doi.org/10.1186/s12887-019-1866-z>

16. Shehab El-Din, E. M. R., El-Sokkary, M. M. A., Basiouny, M. R., Hassan, R. (2015). Epidemiology of Neonatal Sepsis and Implicated Pathogens: A Study from Egypt. *BioMed Research International*, 2015, 1–11. doi: <http://doi.org/10.1155/2015/509484>

17. Shah, B. A., Padbury, J. F. (2014). Neonatal sepsis: an old problem with new insights. *Virulence*, 5 (1), 170–178. doi: <http://doi.org/10.4161/viru.26906>

18. Neonatal infection (early onset): antibiotics for prevention and treatment (2012). *Clinical guideline*. Available at: <https://www.nice.org.uk/guidance/cg149/resources/neonatal-infection-early-onset-antibiotics-for-prevention-and-treatment-pdf-35109579233221>

19. Wynn, J. L., Wong, H. R., Shanley, T. P., Bizzarro, M. J., Saiman, L., Polin, R. A. (2014). Time for a Neonatal-Specific Consensus Definition for Sepsis. *Pediatric Critical Care Medicine*, 15 (6), 523–528. doi: <http://doi.org/10.1097/pcc.0000000000000157>

20. Klingenberg, C., Kornelisse, R. F., Buonocore, G., Maier, R. F., Stocker, M. (2018). Culture-Negative Early-Onset Neonatal Sepsis – At the Crossroad Between Efficient Sepsis Care and Antimicrobial Stewardship. *Frontiers in Pediatrics*, 6. doi: <http://doi.org/10.3389/fped.2018.00285>

21. Kostiuk, O. O., Shunko, Ye. Ye., Krasnova, Yu. Yu. (2014). Rannii neonatalnyi sepsys. Osnovni napriamky diahnozyky ta likuvannia. *Neonatolohiia, khirurgiia ta perynatalna medytsyna*, 4 (3 (13)), 104–109.

22. Dritsakou, K., Liosis, G., Gioni, M., Glynou, E., Avdeliodi, K., Papagaroufalas, K. (2014). CRP levels in extremely low birth weight (ELBW) septic infants. *The Journal of Maternal-Fetal & Neonatal Medicine*, 28 (2), 237–239. doi: <http://doi.org/10.3109/14767058.2014.908842>

23. Hofer, N., Jank, K., Strenger, V., Pansy, J., Resch, B. (2015). Inflammatory indices in meconium aspiration syndrome. *Pediatric Pulmonology*, 51 (6), 601–606. doi: <http://doi.org/10.1002/ppul.23349>

24. Muniraman, H., Gardner, D., Skinner, J., Paweletz, A., Vayalakkad, A., Chee, Y. H. et al. (2017). Biomarkers of hepatic injury and function in neonatal hypoxic ischemic encephalopathy and with therapeutic hypothermia. *European Journal of Pediatrics*, 176 (10), 1295–1303. doi: <http://doi.org/10.1007/s00431-017-2956-2>

25. Lee, J., Bang, Y. H., Lee, E. H., Choi, B. M., Hong, Y. S. (2017). The influencing factors on procalcitonin values in newborns with noninfectious conditions during the first week of life. *Korean Journal of Pediatrics*, 60 (1), 10–16. doi: <http://doi.org/10.3345/kjp.2017.60.1.10>

26. Bellos, I., Fitrou, G., Daskalakis, G., Thomakos, N., Papantoniou, N., Pergialiotis, V. (2018). Soluble TREM-1 as a predic-

tive factor of neonatal sepsis: a meta-analysis. *Inflammation Research*, 67 (7), 571–578. doi: <http://doi.org/10.1007/s00011-018-1149-4>

27. Parri, N., Trippella, G., Lisi, C., De Martino, M., Galli, L., Chiappini, E. (2019). Accuracy of presepsin in neonatal sepsis: systematic review and meta-analysis. *Expert Review of Anti-Infective Therapy*, 17 (4), 223–232. doi: <http://doi.org/10.1080/14787210.2019.1584037>

28. Shunko, Ye. Ye. (Ed.) (2014). *Neonatolohiia: natsionalnyi pidruchnyk*. Vol. 1. Kyiv, 960.

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## ORGAN-SAVING AND RECONSTRUCTIVE PLASTIC SURGERY OF BREAST CANCER

p. 53-57

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*Surgical treatment of breast cancer is an integral part of complex treatment. In recent years, the goal of treatment is not only to ensure long-term relapse-free and metastatic survival, but also a proper quality of life, which requires new approaches to surgery.*

*The aim of the literature review was to analyze current trends in the development of methods of surgical treatment of the breast cancer and ways to improve immediate and late results.*

*The results of the analysis and discussion. Until recently, radical mastectomy was performed to treat breast cancer, which causes long-term lymphovenous complications and significantly*

impairs the quality of life of patients. In recent decades, there has been a development of breast cancer surgery in the direction of the introduction of organ-saving and reconstructive plastic surgery with the improvement of technology for their implementation. There is no generally accepted concept of the optimal method of surgical treatment, both in terms of the volume of the operation and the choice of the method of reconstruction.

**Conclusions.** The problem of choosing the optimal method of surgical treatment of breast cancer, which should provide oncological radicalism and at the same time a good cosmetic result remains unresolved and requires further research

**Keywords:** breast cancer, surgical treatment, organ-saving operations, breast reconstruction, quality of life

## References

1. Global, regional, and national life expectancy, all-cause mortality, and cause-specific mortality for 249 causes of death, 1980–2015: a systematic analysis for the Global Burden of Disease Study 2015 (2016). *Lancet*, 388, 1459–1544. Available at: [http://www.thelancet.com/pdfs/journals/lancet/PIIS0140-6736\(16\)31012-1.pdf](http://www.thelancet.com/pdfs/journals/lancet/PIIS0140-6736(16)31012-1.pdf)
2. Ghoncheh, M., Pournamdar, Z., Salehiniya, H. (2016). Incidence and Mortality and Epidemiology of Breast Cancer in the World. *Asian Pacific Journal of Cancer Prevention*, 17 (sup3), 43–46. doi: <http://doi.org/10.7314/apjcp.2016.17.s3.43>
3. Fedorenko, Z. P., Hulak, L. O., Mykhailovych, Yu. Y., Horokh, Ye. L., Ryzhov, A. Yu., Sumkina, O. V. et. al. (2018). Rak v Ukraini, 2016–2017. *Zakhvoriuvanist, smertnist, pokaznyky diialnosti onkologichnoi sluzhby. Biuletten natsionalnoho kantser-reiestru Ukrainy*. Kyiv, 19. Available at: [http://www.ncru.inf.ua/publications/BULL\\_19/index.htm](http://www.ncru.inf.ua/publications/BULL_19/index.htm)
4. Zikiryakhodzhaev, A. D., Rasskazova, E. A., Tukmakov, A. Y., Shirokikh, I. M. (2019). Relapses after radical subcutaneous/skin-sparing mastectomy with simultaneous reconstruction in breast cancer. *Research'n Practical Medicine Journal*, 6 (1), 33–40. doi: <http://doi.org/10.17709/2409-2231-2019-6-1-3>
5. Holleczeck, B., Stegmaier, C., Radosa, J. C., Solomayer, E.-F., Brenner, H. (2019). Risk of loco-regional recurrence and distant metastases of patients with invasive breast cancer up to ten years after diagnosis – results from a registry-based study from Germany. *BMC Cancer*, 19 (1). doi: <http://doi.org/10.1186/s12885-019-5710-5>
6. Solodkii, V. A., Sherstneva, T. V., Meskikh, E. V., Izmailov, T. R. (2018). The state of the issue of reconstructive plastic surgery for breast cancer in the Russian Federation and in the world. *Vestnik Nacionalnogo mediko-khirurgicheskogo Centra im. N. I. Pirogova*, 13 (3), 132–137. doi: <http://doi.org/10.25881/BPNMSC.2018.78.94.028>
7. Malygin, S. E. (2015). The role of mastectomy in treatment and prophylaxis of breast cancer: beginning, evolution and recent changes. *Malignant Tumours*, 4, 3–13. doi: <http://doi.org/10.18027/2224-5057-2015-4-3-13>
8. Aerts, L., Christiaens, M. R., Enzlin, P., Neven, P., Amant, F. (2014). Sexual functioning in women after mastectomy versus breast conserving therapy for early-stage breast cancer: A prospective controlled study. *The Breast*, 23 (5), 629–636. doi: <http://doi.org/10.1016/j.breast.2014.06.012>
9. Urban, C., Anselmi, K. F., Kuroda, F., Schwartz, J.-C. (2014). Oncoplasty as the Standard of Care in Breast Cancer Surgery. *European Oncology & Haematology*, 10 (1), 43–47. doi: <http://doi.org/10.17925/eoh.2014.10.1.43>
10. Galich, S. P., Pinchuk, V. D. (2011). *Rekonstruktivnaia khirurgiia grudi*. Kyiv: Kniga-plius, 264.
11. Sobolevskii, V. A., Ivashkov, V. Iu., Mekhtieva, N. I. G. (2017). Reconstructive and plastic surgery for breast cancer. *Prakticheskaia onkologiya*, 18 (3), 246–255. Available at: <https://practical-oncology.ru/articles/593.pdf>
12. Watts, G. (2016). Umberto Veronesi. *The Lancet*, 388 (10060), 2600. doi: [http://doi.org/10.1016/s0140-6736\(16\)32230-9](http://doi.org/10.1016/s0140-6736(16)32230-9)
13. Zikiriakhodzhaev, A., Rasskazova, E. (2015). Organ-sparing surgery for breast cancer. *Vrach*, 6, 40–42. Available at: <http://onco.rusvrach.ru/archive/25877305-2015-06-13.pdf>
14. Maxwell, G. P., Storm-Dickerson, T., Whitworth, P., Rubano, C., Gabriel, A. (2011). Advances in Nipple-Sparing Mastectomy: Oncological Safety and Incision Selection. *Aesthetic Surgery Journal*, 31 (3), 310–319. doi: <http://doi.org/10.1177/1090820x11398111>
15. Kinoshita, S., Kyoda, S., Hirano, A., Akiba, T., Nojima, K., Uchida, K. et. al. (2013). Clinical comparison of four types of skin incisions for skin-sparing mastectomy and immediate breast reconstruction. *Surgery Today*, 44 (8), 1470–1475. doi: <http://doi.org/10.1007/s00595-013-0722-2>
16. Lai, H.-W., Wu, H.-S., Chuang, K.-L., Chen, D.-R., Chang, T.-W., Kuo, S.-J. et. al. (2014). Endoscopy-assisted total mastectomy followed by immediate pedicled transverse rectus abdominis musculocutaneous (TRAM) flap reconstruction: preliminary results of 48 patients. *Surgical Innovation*, 22 (4), 382–389. doi: <http://doi.org/10.1177/1553350614546003>
17. Lai, H.-W., Chen, S.-T., Chen, D.-R., Chen, S.-L., Chang, T.-W., Kuo, S.-J. et. al. (2016). Current Trends in and Indications for Endoscopy-Assisted Breast Surgery for Breast Cancer: Results from a Six-Year Study Conducted by the Taiwan Endoscopic Breast Surgery Cooperative Group. *PLOS ONE*, 11(3), e0150310. doi: [10.1371/journal.pone.0150310](https://doi.org/10.1371/journal.pone.0150310)
18. Tukenmez, M., Ozden, B. C., Agcaoglu, O., Kecer, M., Ozmen, V., Muslumanoglu, M., Igci, A. (2014). Videoendoscopic Single-Port Nipple-Sparing Mastectomy and Immediate Reconstruction. *Journal of Laparoendoscopic & Advanced Surgical Techniques*, 24 (2), 77–82. doi: <http://doi.org/10.1089/lap.2013.0172>
19. Bit-Sava, E. M., Egorenkov, V. V., Damenia, A. O., Melnikova, O. A., Akhmedov, R. M., Monogorova, M. A. et. al. (2017). New approaches in breast cancer surgery. *Practical Oncology*, 18 (3), 232–245. doi: <http://doi.org/10.31917/1803232>
20. Ermoshchenkova, M. V., Chissov, V. I., Usov, A. V., Sukhotko, A. S., Tukmakov, A. Y., Baichorov, E. A., Zikiryahodjaev, A. D. (2017). Biological and synthetic mesh use in reconstructive surgery in patients with breast cancer. *Research'n Practical Medicine Journal*, 4 (1), 23–32. doi: <http://doi.org/10.17709/2409-2231-2017-4-1-3>
21. Franceschini, G., Martin, S. A., Di Leon, A., Magno, A., Moschella, F., Accetta, C., Masetti, R. (2015). New trends in breast cancer surgery: a therapeutic approach increasingly efficacy and respectful of the patient. *Giornale di Chirurgia – Journal of Surgery*, 36 (4), 145–152. doi: <http://doi.org/10.11138/gchir/2015.36.4.145>
22. Hernanz, F., Sánchez, S., Cerdeira, M. P., Figuero, C. R. (2011). Long-term results of breast conservation and immediate volume replacement with myocutaneous latissimus dorsi flap. *World Journal of Surgical Oncology*, 9 (1). doi: <http://doi.org/10.1186/1477-7819-9-159>
23. Wang, X.-L., Liu, L.-B., Song, F.-M., Wang, Q.-Y. (2014). Meta-analysis of the Safety and Factors Contributing to Complications of MS-TRAM, DIEP, and SIEA Flaps for Breast Reconstruction. *Aesthetic Plastic Surgery*, 38 (4), 681–691. doi: <http://doi.org/10.1007/s00266-014-0333-3>
24. Simonacci, F., Bertozzi, N., Grieco, M. P., Grignaffini, E., Rapisio, E. (2016). Autologous fat transplantation for breast reconstruction: A literature review. *Annals of Medicine and Surgery*, 12, 94–100. doi: <http://doi.org/10.1016/j.amsu.2016.11.012>

25. Van Turnhout, A. A., Fuchs, S., Lisabeth-Broné, K., Vriens-Nieuwenhuis, E. J. C., van der Sluis, W. B. (2017). Surgical Outcome and Cosmetic Results of Autologous Fat Grafting After Breast Conserving Surgery and Radiotherapy for Breast Cancer: A Retrospective Cohort Study of 222 Fat Grafting Sessions in 109 Patients. *Aesthetic Plastic Surgery*, 41 (6), 1334–1341. doi: <http://doi.org/10.1007/s00266-017-0946-4>
26. Zikiryakhodzhaev, A. D., Efanov, V. V., Usov, F. N., Ismailova, M. M. (2015). Complications of delayed reconstructive and plastic surgery after radical mastectomy. *Tumors of Female Reproductive System*, 11 (2), 31–34. doi: <http://doi.org/10.17650/1994-4098-2015-11-2-31-34>
27. Maxwell, G. P., Gabriel, M. (2017). Breast implant design. *Gland Surgery*, 6 (2), 148–153. doi: <http://doi.org/10.21037/g.2016.11.09>
28. Zikiryakhodzhaev, A. D., Shirokikh, I. M., Ablitsova, N. V., Ermoshchenkova, M. V., Kharchenko, N. V., Saribekyan, E. K. et. al. (2018). Biological and synthetic materials in reconstructive surgery for breast cancer treatment (literature review). *Tumors of Female Reproductive System*, 14 (1), 28–37. doi: <http://doi.org/10.17650/1994-4098-2018-14-1-28-37>
29. Dragun, A. E., Huang, B., Tucker, T. C., Spanos, W. J. (2012). Increasing Mastectomy Rates Among all Age Groups for Early Stage Breast Cancer: A 10-Year Study of Surgical Choice. *The Breast Journal*, 18 (4), 318–325. doi: <http://doi.org/10.1111/j.1524-4741.2012.01245.x>
30. Ho, A. L., Bovill, E. S., Macadam, S. A., Tyldesley, S., Giang, J., Lennox, P. A. (2014). Postmastectomy Radiation Therapy after Immediate Two-Stage Tissue Expander/Implant Breast Reconstruction. *Plastic and Reconstructive Surgery*, 134 (1), 1e–10e. doi: <http://doi.org/10.1097/prs.0000000000000292>
31. Khailenko, D. V., Egorov, Iu. S., Portnoi, S. M., Khailenko, V. A. (2016) Primary reconstructive plastic surgery in nodular form of III stage breast cancer. *Rossiiskii onkologicheskii zhurnal*, 21 (4), 175–178.
32. Cordeiro, P. G., Albornoz, C. R., McCormick, B., Hu, Q., Van Zee, K. (2014). The impact of postmastectomy radiotherapy on two-stage implant breast reconstruction: an analysis of long-term surgical outcomes, aesthetic results, and satisfaction over 13 years. *Plastic and Reconstructive Surgery*, 134 (4), 588–595. doi: <http://doi.org/10.1097/prs.0000000000000523>
33. Nahabedian, M. Y., Jacobson, S. R. (2019). Two-stage prepectoral breast reconstruction. *Gland Surgery*, 8 (1), 43–52. doi: <http://doi.org/10.21037/g.2018.09.04>
34. Casella, D., Di Taranto, G., Marcasciano, M., Lo Torro, F., Barellini, L., Sordi, S. et. al. (2019). Subcutaneous expanders and synthetic mesh for breast reconstruction: Long-term and patient-reported BREAST-Q outcomes of a single-center prospective study. *Journal of Plastic, Reconstructive & Aesthetic Surgery*, 72 (5), 805–812. doi: <http://doi.org/10.1016/j.bjps.2018.12.018>
35. Hon, H. H., Mubang, R. N., Wernick, B. D., Freedman, S. F., Stoltzfus, J. C., Miele, L. F. et. al. (2017). Acellular dermal matrix versus inferior deepithelialized flap breast reconstruction: equivalent outcomes, with increased cost. *Plastic and Reconstructive Surgery – Global Open*, 5 (6), e1382. doi: <http://doi.org/10.1097/gox.0000000000001382>
36. Sorkin, M., Qi, J., Kim, H. M., Hamill, J. B., Kozlow, J. H., Pusic, A. L., Wilkins, E. G. (2017). Acellular dermal matrix in immediate expander/implant breast reconstruction: a multicenter assessment of risks and benefits. *Plastic and Reconstructive Surgery*, 140 (6), 1091–1100. doi: <http://doi.org/10.1097/prs.0000000000003842>
37. Zikiryakhodzhaev, A. D., Rasskazova, E. A. (2018). Two-stage delayed reconstruction in patients with breast cancer after mastectomy. *Malignant Tumours*, 8 (2), 43–49. doi: <http://doi.org/10.18027/2224-5057-2018-8-2-43-49>
38. Hvilsum, G. B., Hölmich, L. R., Frederiksen, K., Steinding-Jessen, M., Friis, S., Dalton, S. O. (2010). Socioeconomic position and breast reconstruction in Danish women. *Acta Oncologica*, 50 (2), 265–273. doi: <http://doi.org/10.3109/0284186x.2010.529823>
39. Thiruchelvam, P. T. R., McNeill, F., Jallali, N., Harris, P., Hogben, K. (2013). Post-mastectomy breast reconstruction. *BMJ*, 347 (2), f5903–f5903. doi: <http://doi.org/10.1136/bmj.f5903>
40. Park, E. H., Min, S. Y., Kim, Z., Yoon, C. S., Jung, K.-W. et. al. (2017). Basic Facts of Breast Cancer in Korea in 2014: The 10-Year Overall Survival Progress. *Journal of Breast Cancer*, 20 (1), 1. doi: <http://doi.org/10.4048/jbc.2017.20.1.1>
41. Nguyen, T. T., Hoskin, T. L., Habermann, E. B., Chevillat, A. L., Boughey, J. C. (2017). Breast Cancer-Related Lymphedema Risk is Related to Multidisciplinary Treatment and Not Surgery Alone: Results from a Large Cohort Study. *Annals of Surgical Oncology*, 24 (10), 2972–2980. doi: <http://doi.org/10.1245/s10434-017-5960-x>
42. Madsen, R. J., Esmonde, N. O., Ramsey, K. L., Hansen, J. E. (2016). Axillary Lymph Node Dissection Is a Risk Factor for Major Complications After Immediate Breast Reconstruction. *Annals of Plastic Surgery*, 77 (5), 513–516. doi: <http://doi.org/10.1097/sap.0000000000000653>
43. Marinescu, S. A., Bejinariu, C. G., Şapte, E., Marinaş, M. C., Giuglea, C. (2019). Complications related to breast reconstruction after mastectomy using multiple surgical techniques – a national and international comparative analysis. *Romanian Journal of Morphology and Embryology*, 60 (1), 87–93.
44. Gambardella, C., Clarizia, G., Patrone, R., Offi, C., Mauriello, C., Romano, R. et. al. (2019). Advanced hemostasis in axillary lymph node dissection for locally advanced breast cancer: new technology devices compared in the prevention of seroma formation. *BMC Surgery*, 18 (S1). doi: <http://doi.org/10.1186/s12893-018-0454-8>
45. Vinnyk, Y. O., Vlasenko, V. G., Baranova, A. V. (2019). Prophylaxis of complications after radical operations in patients, suffering a mammary gland cancer. *Klinicheskaia Khirurgiia*, 86 (11-12), 46–50. doi: <http://doi.org/10.26779/2522-1396.2019.11-12.46>