Effect Of Excess Body Weight And Deficiency Of Vitamin A D On The Clinical Course Of Uterine Myoma

Dinara Kh. Irnazarova¹, Dilchekhra K. Yuldasheva², Dilbar K. Najmutdinova³, Dilrabo T. Kayumova⁴, Fotima A. Atakhodjaeva⁵, Gulmira A. Akhmedova⁶, Dilfuza R. Sadikova⁷

¹Irnazarova Dinara Khamidiloyevna MD, PhD doctoral student of Tashkent Medical Academy

²Yuldasheva Dilchekhra Yusupkhonovna MD, PhD, DSc, Associate Professor of the Department of Obstetrics and Gynecology No. 2 TMA

³Najmutdinova Dilbar Kamaritdinovna d . m . n ., Professor , Head of Department of Obstetrics and Gynecology №2 TMA

⁴Dilrabo Talmasovna Kayumova , a PhD, to . m . n ., Associate Professor, Department of Obstetrics and Gynecology №2 TMA

⁵Atakhodjayeva Fotima Abduraimovna to . m . n ., Associate Professor, Department of Obstetrics and Gynecology №2 TMA

⁶Akhmedova Gulmira Amanovna, to . m . n ., Assistant Associate Professor of the Department of Obstetrics and Gynecology №2 TMA

⁷Sadikova Dilfuza Ravshanbekovna MD, PhD, Associate Professor of the Department of Obstetrics and Gynecology No. 2 TMA

Email: ¹dinarairnazarova@mail.ru, ²dilchehra@list.ru, ³dibarkn@mail.ru, ⁴kdilrabo@mail.ru, ⁵fatima.atakhodjaeva@tma.uz, ⁶gulmura.akhmedova@tma.uz, ⁷dr.sadikova1974@gmail.com

ABSTRACT

Uterine fibroids is the most common benign gynecological disease, the pathogenesis of which is not fully understood. We examined 152 women in the II clinic of the T ashkent Medical Academy. The saturation of the body with vitamin D (25 (OH)D) and its relationship with concomitant obesity in patients with uterine fibroids and healthy women were studied. In less than half of women with symptomatic myoma, deficiency and severe deficiency of vitamin D prevails with concomitant obesity of varying degrees. Studying the relationship between vitamin D deficiency and excess body weight in this pathology makes it possible to predict the development of the disease, the nature of the course and the risk of possible complications.

Keywords: uterine fibroids, vitamin D (prohormone D), risk factors, body mass index (BMI), obesity.

1. INTRODUCTION

Leiomyomas are benign clonal tumors in women [11]. M iomu uterus diagnosed in 30-35% of women of reproductive age, usually in late reproductive age [16], and in 1/3 of patients it becomes symptomatic [1, 18, 20], and by the age of 50- 80% of women [17, 18].

The authors describe risk factors for the development of fibroids (age before menopause, black race, obesity), reproductive (infertility, earlier menarche, use of oral contraception under 16 years of age, etc.) and environmental (diet, reduced insolation, leading to vitamin D deficiency, environmental toxins environment) factors that are the subject of current research [12, 13]. Given the high prevalence of the disease in the population, at the present stage, the study of risk factors in the development of the disease is one of the topical, but, unfortunately, not fully studied issues in gynecology and is still a subject of discussion. According to many researchers, obesity is a significant potentiating endocrine factor of the disease and occurs in women with fibroids higher in 25-70% of cases [2].

Several studies show that vitamin D deficiency is a risk factor for the development of uterine fibroids [8,9,12,19,21,23]. The protective role of vitamin D on the growth of fibroids has been demonstrated during in vitro [10] and in vivo [8] studies, as well as retrospective clinical studies [3], which indicate the existence of a clear protective effect of vitamin D on the growth of fibroids, which is based on lie inhibition of cell proliferation, stimulation of apoptosis and other pharmacodynamic effects. The next logical step would be to demonstrate the inhibitory effect of vitamin D in humans, which would require clinical research. The authors argue that vitamin D deficiency is associated with fibroids and that supplementation helps nodule shrinkage and disease regression [14,25].

The aim of the study is to identify the level of vitamin D supply and the degree of obesity, and their relationship with the clinical course in women with uterine fibroids.

2. MATERIALS AND RESEARCH METHODS

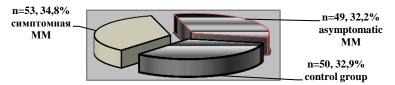
The design of a cohort prospective controlled study was used. The study was based on clinical and laboratory examination of 152 women who were admitted to the 2nd clinic of the Tashkent Medical Academy. The surveyed women were divided into 2 groups: the control group consisted of 50 healthy women and the main group of 102 women with uterine myoma. Main group patients were subdivided into two subgraphs uppy - 53 women with symptomatic iomoy and 49 women - asymptomatic . [15].

During the observation period, all women underwent identification of risk factors according to a modified scale recommended by the guidelines, general clinical examination, ultrasound of the uterus and appendages with duplex scanning of the uterine artery, morphological studies of aspirates from the uterus. The marker of the body saturation with vitamin D - 25 (OH) D was determined by the method of ELISA quantitative determination - chemiluminescence analysis on microparticles (CMIA) [7]. Mathematical processing and statistical analysis of the results obtained were carried out using the program "Statistika 6.0".

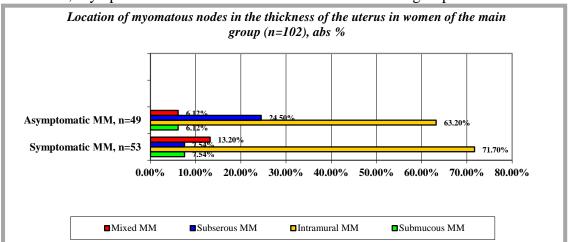
3. RESEARCH RESULTS AND THEIR DISCUSSION

The age of the surveyed women in the main group was 19-55 years, the average age in the first subgroup was 44.35 ± 0.83 (n = 53) and in the second subgroup 42.6 ± 0.7 years (n = 49), which corresponds to the authors' data [6, 16, 22] that fibroids are more often diagnosed in the late reproductive period. Whereas in the control group (n = 50), the average age was 40.12 ± 0.7 years (p < 0.01).

Division of observed women into groups n, abs%



Ultrasound study on the myoma node localization in the thickness of the uterus in both groups the main group prevailed intramural node (71.7% and 63.2%, respectively groups. Mixed fibroids in women with symptomatic fibroids were 2 times more likely than women with asymptomatic myoma (13.2% and 6.12%, respectively). The median uterine volume calculated according to Brunn's formula (1981) in the subgroup with symptomatic MM was 237.54 mm 3 , asymptomatic MM - 103.45 mm 3 and in the control group - 52.1 m m 3 .

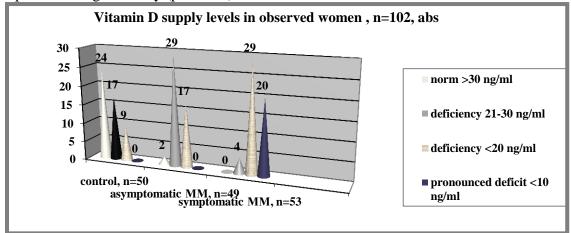


By analysis of risk factors prevalent factors such as overweight BMI (29.7 \pm 11.83 and 28.1 \pm 0.08 kg / m 2 , respectively subgroups and the control group of 23.3 \pm 0.01 kg / m 2 , p <0.01), burdened obstetric and gynecological history. According to WHO (2012), vitamin D levels are affected by dietary intake of vitamin D and factors affecting its absorption metabolism, as well as obesity. Researchers have described the relationship between excess BMI and vitamin D deficiency [9, 24]. Our analysis to identify excess BMI showed that in 47.2% of women with symptomatic MM, obesity of I, II, III degrees was calculated (28.3%, 11.3% and 7.5%, respectively) [15].

Women with symptomatic myoma of the main group (n=53) applied to various clinics: the symptom of bleeding and anemia prevailed to a greater extent in 83.01% (n=44), of which 16.9% underwent blood transfusion due to severe anemia; a symptom of rapid growth - 9.43% (n=5), a symptom of infertility in 5.6% (n=3) and a symptom of pelvic pain (n=2) 3.77%.

The status of vitamin D in the studied women showed that the values in the group of women with fibroids ranged from 4 to 36 ng / ml and averaged 16.7 ± 1.8 ng / ml, which turned out to be significantly lower than in healthy women (p < 0.001). It is important to note that, according to the researchers, the determination of the initial serum level of prohormone 25 (OH) D by laboratory methods is the most acceptable, reliable and clinically significant for assessing the saturation of vitamin D in the human body [5]. When assessing the content of vitamin D in the blood in the structure of the main group in women with symptomatic MM, the average was 11.84 ± 0.46 ng / ml and in asymptomatic - 21.54 ± 0.04 , while in the control group 29, 83 ± 1.13 ng / ml (p <0.001). At the same time, a pronounced deficiency of

prohormone D was detected in the subgroup of symptomatic MM 6.62 ± 0.9 ng / ml in 37.7% of women with an obvious clinical manifestation of menorrhagia (in 100%) and a recurrent course of the disease, and in the subgroup of asymptomatic myoma in 1 / 3 patients (36.7%) have a vitamin D deficiency 16.7 ± 1.6 [4]. The prohormone values in the control group differed significantly (p <0.001).



When comparing vitamin D indices between the subgroups of the main group, the statistical differences were significant, which indicated the presence of a relationship between the level of vitamin D saturation in women with MM and clinical manifestations of the disease and the size of the uterus. The distribution of women according to the degree of vitamin D provision, based on its content in the blood [15], showed a significant difference between the subgroups of the main group and healthy ones.

No.	Groups	Symptomatic MM, n =		Asymptomatic MM, n		Control group, n =	
		53		= 49		50	
	Criteria	n, abs	(ng/ml)	n, abs	(ng/ml)	n, abs	(ng/ml)
1	Norm	0	-	2	34 <u>+</u> 1.2 ''	24	40.4 <u>+</u> 1.7
				(4.08%)	_	(48%)	_
2	Failure	4	20.8 <u>+</u> 1.9	29	23.6 <u>+</u> 1.4 ''	17	24.4 <u>+</u> 1.7
		(7.54%)	***	(59.2%)	_	(34%)	_
3	Deficit	29	14.02 <u>+</u> 0.2	17	16.8 <u>+</u> 1.6 "	9	14.2 <u>+</u> 0.9
		(54.7%)	***	(34.7%)		(18%)	
4	Severe deficiency	20	6.62 <u>+</u> 0.9	0	-	0	-
		(37.7%)	***				

Note: * p < 0.001 in relation to the first subgroup; - 'p < 0.001 with respect to the second subgroup.

Table 1. Values of vitamin D levels in women in the study groups.

In the main group of women with asymptomatic MM, normal values of vitamin D were only 4.08% of cases, insufficiency - in more than half (59.2%) and deficiency - in 1/3 of women, in the subgroup of symptomatic fibroids, normal values were not found in women, deficit - 54.7% and pronounced deficit - 37.7%, which is 2.88 times more than in the group of asymptomatic myoma and 5.4 times more than in the control group. It is noteworthy that, despite the absence of the disease, 52% of healthy women had an insufficient level and deficiency of vitamin D in the blood, in a country with sufficient insolation. Correlation analysis showed a direct weak positive relationship in the control group between the content of vitamin D in the blood and BMI (r = 0.345, p < 0.001), that is, normal values of vitamin D corresponded to a normal level of BMI; while the correlation between these indicators in the

main group, especially in the subgroup of symptomatic myoma, was a direct mean positive (r = 0.482, p < 0.001).

When analyzing the odds ratio (OR) of MM in women with deficiency and / or deficiency of vitamin D, it was revealed: in women with insufficiency and / or deficiency of vitamin D, the chances of developing MM is OR = 16.13, which proves that the factor of prohormone deficiency is strong a provoking factor for the development of the disease in these women (Table 2).

Odds ratio (OR)	Study	women	with	Study women without
	MM			MM
Vitamin D deficiency /	100			74
deficiency				
Vitamin D norm	2			24
Total	102			98

100/2 = 5074/24 = 3.1

50/3.1 = 16.13 (OR) positive strong factor

Table 2. Odds ratio (OR) of developing MM in women with vitamin D deficiency / deficiency.

When analyzing the odds ratio (OR) of MM in women with obesity, it was revealed that in obese women the chances of developing MM are OR = 7.38, which proves that excess weight is a factor promoter for the development of severe clinical symptoms of the disease in these women, reducing the quality life of women of this cohort (Table 3).

Odds ratio (OR)	Study women with MM	Study	women	without
		MM		
Women with excess BMI and	78	thirty		
obesity				
Women without obesity	24	68		
Total	102	98		

 $78/24 = 3.25 \ 30/68 = 0.44$

3.25 / 0.44 = 7.38 (OR) positive strong factor

Table 3. Odds ratio (OR) of developing MM in women with excess BMI and obesity.

4. DISCUSSION

Our results showed that lower values of vitamin D in patients with fibroids with concomitant excess BMI and obesity can affect the nature of the course of the disease and the incidence of complications. N atsientki with severe deficiency of vitamin D in the subgroup symptomatic uterine myoma (<10 ng / ml) had a bright clinical manifestations of the disease: menorrhagia, leading to anemizatsii women in this category, the rapid growth myoma node amid critical deficiency of vitamin D and obesity, accompanying with pelvic pain, infertility m. In women with symptomatic myoma, the level of vitamin D was significantly lower in the group compared with the group with asymptomatic myoma.

5. CONCLUSION

A positive correlation between blood vitamin D and elevated BMI is especially noted in the subgroup of symptomatic uterine fibroids. D cial studies to detail the mechanisms of the effect of vitamin D and the modifying factors - obesity on for uterine fibroids may in the future find widespread use nenie health and become the basis of modern second personalized medicine, because it gives the opportunity to predict the development of the disease, the nature of the flow and the risk of possible complications.

List of references:

- [1] Kant, N., Saralch, S., & Singh, H. (2011). Ponderomotive self-focusing of a short laser pulse under a plasma density ramp. *Nukleonika*, *56*, 149-153.
- [2] Patyar, S., & Patyar, R. R. (2015). Correlation between sleep duration and risk of stroke. *Journal of Stroke and Cerebrovascular Diseases*, 24(5), 905-911.
- [3] Khamparia, A., & Pandey, B. (2015). Knowledge and intelligent computing methods in e-learning. *International Journal of technology enhanced learning*, 7(3), 221-242.
- [4] Singh, A., Lin, Y., Quraishi, M. A., Olasunkanmi, L. O., Fayemi, O. E., Sasikumar, Y., ... & Kabanda, M. M. (2015). Porphyrins as corrosion inhibitors for N80 Steel in 3.5% NaCl solution: Electrochemical, quantum chemical, QSAR and Monte Carlo simulations studies. *Molecules*, 20(8), 15122-15146.
- [5] Singh, S., Kumar, V., Upadhyay, N., Singh, J., Singla, S., & Datta, S. (2017). Efficient biodegradation of acephate by Pseudomonas pseudoalcaligenes PS-5 in the presence and absence of heavy metal ions [Cu (II) and Fe (III)], and humic acid. *3 Biotech*, 7(4), 262.
- [6] Mia, M., Singh, G., Gupta, M. K., & Sharma, V. S. (2018). Influence of Ranque-Hilsch vortex tube and nitrogen gas assisted MQL in precision turning of Al 6061-T6. *Precision Engineering*, 53, 289-299.
- [7] Prakash, C., Singh, S., Pabla, B. S., & Uddin, M. S. (2018). Synthesis, characterization, corrosion and bioactivity investigation of nano-HA coating deposited on biodegradable Mg-Zn-Mn alloy. *Surface and Coatings Technology*, *346*, 9-18.
- [8] Feng, X., Sureda, A., Jafari, S., Memariani, Z., Tewari, D., Annunziata, G., ... & Sychrová, A. (2019). Berberine in cardiovascular and metabolic diseases: from mechanisms to therapeutics. *Theranostics*, 9(7), 1923.
- [9] Bashir, S., Sharma, V., Lgaz, H., Chung, I. M., Singh, A., & Kumar, A. (2018). The inhibition action of analgin on the corrosion of mild steel in acidic medium: A combined theoretical and experimental approach. *Journal of Molecular Liquids*, 263, 454-462.
- [10] Sidhu, G. K., Singh, S., Kumar, V., Dhanjal, D. S., Datta, S., & Singh, J. (2019). Toxicity, monitoring and biodegradation of organophosphate pesticides: a review. *Critical Reviews in Environmental Science and Technology*, 49(13), 1135-1187.
- [11] Nanda, V., & Kant, N. (2014). Enhanced relativistic self-focusing of Hermite-cosh-Gaussian laser beam in plasma under density transition. *Physics of Plasmas*, 21(4), 042101.
- [12] Kotla, N. G., Gulati, M., Singh, S. K., & Shivapooja, A. (2014). Facts, fallacies and future of dissolution testing of polysaccharide based colon-specific drug delivery. *Journal of Controlled Release*, 178, 55-62.
- [13] Farooq, R., & Shankar, R. (2016). Role of structural equation modeling in scale development. *Journal of Advances in Management Research*.
- [14] Singh, S., Ramakrishna, S., & Gupta, M. K. (2017). Towards zero waste manufacturing: A multidisciplinary review. *Journal of cleaner production*, *168*, 1230-1243.
- [15] Mahla, S. K., Dhir, A., Gill, K. J., Cho, H. M., Lim, H. C., & Chauhan, B. S. (2018). Influence of EGR on the simultaneous reduction of NOx-smoke emissions trade-off under CNG-biodiesel dual fuel engine. *Energy*, 152, 303-312.
- [16] Nanda, V., Kant, N., & Wani, M. A. (2013). Self-focusing of a Hermite-cosh Gaussian laser beam in a magnetoplasma with ramp density profile. *Physics of Plasmas*, 20(11), 113109.

- [17] Kaur, P., Singh, S. K., Garg, V., Gulati, M., & Vaidya, Y. (2015). Optimization of spray drying process for formulation of solid dispersion containing polypeptide-k powder through quality by design approach. *Powder Technology*, 284, 1-11.
- [18] Sharma, D., & Saharan, B. S. (2016). Functional characterization of biomedical potential of biosurfactant produced by Lactobacillus helveticus. *Biotechnology Reports*, 11, 27-35.
- [19] Wani, A. B., Chadar, H., Wani, A. H., Singh, S., & Upadhyay, N. (2017). Salicylic acid to decrease plant stress. *Environmental Chemistry Letters*, 15(1), 101-123.
- [20] Mishra, V., Patil, A., Thakur, S., & Kesharwani, P. (2018). Carbon dots: emerging theranostic nanoarchitectures. *Drug discovery today*, 23(6), 1219-1232.
- [21] Kumar, V., Pitale, S. S., Mishra, V., Nagpure, I. M., Biggs, M. M., Ntwaeaborwa, O. M., & Swart, H. C. (2010). Luminescence investigations of Ce3+ doped CaS nanophosphors. *Journal of alloys and compounds*, 492(1-2), L8-L12.
- [22] Pudake, R. N., Swaminathan, S., Sahu, B. B., Leandro, L. F., & Bhattacharyya, M. K. (2013). Investigation of the Fusariumvirguliformefvtox1 mutants revealed that the FvTox1 toxin is involved in foliar sudden death syndrome development in soybean. *Current genetics*, 59(3), 107-117.
- [23] Kapoor, B., Singh, S. K., Gulati, M., Gupta, R., & Vaidya, Y. (2014). Application of liposomes in treatment of rheumatoid arthritis: quo vadis. *The scientific world Journal*, 2014.
- [24] Haldhar, R., Prasad, D., & Saxena, A. (2018). Myristica fragrans extract as an ecofriendly corrosion inhibitor for mild steel in 0.5 M H2SO4 solution. *Journal of Environmental Chemical Engineering*, 6(2), 2290-2301.
- [25] Bordoloi, N., Sharma, A., Nautiyal, H., & Goel, V. (2018). An intense review on the latest advancements of Earth Air Heat Exchangers. *Renewable and Sustainable Energy Reviews*, 89, 261-280.
- [26] Sharma, P., Mehta, M., Dhanjal, D. S., Kaur, S., Gupta, G., Singh, H., ... & Chellappan, D. K. (2019). Emerging trends in the novel drug delivery approaches for the treatment of lung cancer. *Chemico-biological interactions*, 309, 108720.
- [27] Goga, G., Chauhan, B. S., Mahla, S. K., & Cho, H. M. (2019). Performance and emission characteristics of diesel engine fueled with rice bran biodiesel and n-butanol. *Energy Reports*, 5, 78-83.
- [28] Umashankar, M. S., Sachdeva, R. K., & Gulati, M. (2010). Aquasomes: a promising carrier for peptides and protein delivery. *Nanomedicine: Nanotechnology, Biology and Medicine*, 6(3), 419-426.
- [29] Sharma, A., Shree, V., & Nautiyal, H. (2012). Life cycle environmental assessment of an educational building in Northern India: A case study. *Sustainable Cities and Society*, 4, 22-28.
- [30] Kaur, T., Kumar, S., Bhat, B. H., Want, B., & Srivastava, A. K. (2015). Effect on dielectric, magnetic, optical and structural properties of Nd–Co substituted barium hexaferrite nanoparticles. *Applied Physics A*, 119(4), 1531-1540.
- [31] Datta, S., Singh, J., Singh, S., & Singh, J. (2016). Earthworms, pesticides and sustainable agriculture: a review. *Environmental Science and Pollution Research*, 23(9), 8227-8243.
- [32] Vij, S., & Bedi, H. S. (2016). Are subjective business performance measures justified?. *International Journal of Productivity and Performance Management*.
- [33] Chawla, R., & Sharma, S. (2017). Molecular dynamics simulation of carbon nanotube pull-out from polyethylene matrix. *Composites Science and Technology*, 144, 169-177.

- [34] Prakash, C., & Uddin, M. S. (2017). Surface modification of β-phase Ti implant by hydroaxyapatite mixed electric discharge machining to enhance the corrosion resistance and in-vitro bioactivity. *Surface and Coatings Technology*, 326, 134-145.
- [35] Saxena, A., Prasad, D., & Haldhar, R. (2018). Investigation of corrosion inhibition effect and adsorption activities of Cuscuta reflexa extract for mild steel in 0.5 M H2SO4. *Bioelectrochemistry*, 124, 156-164.
- [36] Prabhakar, P. K., Kumar, A., & Doble, M. (2014). Combination therapy: a new strategy to manage diabetes and its complications. *Phytomedicine*, 21(2), 123-130.
- [37] Wheeler, K. C., Jena, M. K., Pradhan, B. S., Nayak, N., Das, S., Hsu, C. D., ... & Nayak, N. R. (2018). VEGF may contribute to macrophage recruitment and M2 polarization in the decidua. *PLoS One*, *13*(1), e0191040.
- [38] Singh, A., Lin, Y., Ansari, K. R., Quraishi, M. A., Ebenso, E. E., Chen, S., & Liu, W. (2015). Electrochemical and surface studies of some Porphines as corrosion inhibitor for J55 steel in sweet corrosion environment. *Applied Surface Science*, 359, 331-339.
- [39] Gill, J. P. K., Sethi, N., Mohan, A., Datta, S., & Girdhar, M. (2018). Glyphosate toxicity for animals. *Environmental Chemistry Letters*, 16(2), 401-426.
- [40] Kumar, V., Singh, S., Singh, J., & Upadhyay, N. (2015). Potential of plant growth promoting traits by bacteria isolated from heavy metal contaminated soils. *Bulletin of environmental contamination and toxicology*, 94(6), 807-814.
- [41] Patel, S. (2012). Potential of fruit and vegetable wastes as novel biosorbents: summarizing the recent studies. *Reviews in Environmental Science and Bio/Technology*, 11(4), 365-380.
- [42] Srivastava, G., Das, C. K., Das, A., Singh, S. K., Roy, M., Kim, H., ... & Philip, D. (2014). Seed treatment with iron pyrite (FeS 2) nanoparticles increases the production of spinach. *RSC Advances*, 4(102), 58495-58504.
- [43] Nagpal, R., Behare, P. V., Kumar, M., Mohania, D., Yadav, M., Jain, S., ... & Henry, C. J. K. (2012). Milk, milk products, and disease free health: an updated overview. *Critical reviews in food science and nutrition*, 52(4), 321-333.
- [44] Vaid, S. K., Kumar, B., Sharma, A., Shukla, A. K., & Srivastava, P. C. (2014). Effect of Zn solubilizing bacteria on growth promotion and Zn nutrition of rice. *Journal of soil science and plant nutrition*, *14*(4), 889-910.
- [45] Lin, Y., Singh, A., Ebenso, E. E., Wu, Y., Zhu, C., & Zhu, H. (2015). Effect of poly (methyl methacrylate-co-N-vinyl-2-pyrrolidone) polymer on J55 steel corrosion in 3.5% NaCl solution saturated with CO2. *Journal of the Taiwan Institute of Chemical Engineers*, 46, 214-222.
- [46] Mahesh, K. V., Singh, S. K., & Gulati, M. (2014). A comparative study of top-down and bottom-up approaches for the preparation of nanosuspensions of glipizide. *Powder technology*, 256, 436-449.
- [47] Singh, G., Gupta, M. K., Mia, M., & Sharma, V. S. (2018). Modeling and optimization of tool wear in MQL-assisted milling of Inconel 718 superalloy using evolutionary techniques. *The International Journal of Advanced Manufacturing Technology*, 97(1-4), 481-494.
- [48] Chauhan, C. C., Kagdi, A. R., Jotania, R. B., Upadhyay, A., Sandhu, C. S., Shirsath, S. E., & Meena, S. S. (2018). Structural, magnetic and dielectric properties of Co-Zr substituted M-type calcium hexagonal ferrite nanoparticles in the presence of α-Fe2O3 phase. *Ceramics International*, 44(15), 17812-17823.
- [49] Sharma, A., Shahzad, B., Kumar, V., Kohli, S. K., Sidhu, G. P. S., Bali, A. S., ... & Zheng, B. (2019). Phytohormones regulate accumulation of osmolytes under abiotic stress. *Biomolecules*, 9(7), 285.

- [50] Balakumar, P., Chakkarwar, V. A., Kumar, V., Jain, A., Reddy, J., & Singh, M. (2008). Experimental models for nephropathy. *Journal of the Renin-Angiotensin-Aldosterone System*, 9(4), 189-195.
- [51] Singh, A., Lin, Y., Liu, W., Kuanhai, D., Pan, J., Huang, B., ... & Zeng, D. (2014). A study on the inhibition of N80 steel in 3.5% NaCl solution saturated with CO2 by fruit extract of Gingko biloba. *Journal of the Taiwan Institute of Chemical Engineers*, 45(4), 1918-1926.
- [52] Kaur, T., Kaur, B., Bhat, B. H., Kumar, S., & Srivastava, A. K. (2015). Effect of calcination temperature on microstructure, dielectric, magnetic and optical properties of Ba0. 7La0. 3Fe11. 7Co0. 3O19 hexaferrites. *Physica B: Condensed Matter*, 456, 206-212
- [53] Singh, P., Singh, A., & Quraishi, M. A. (2016). Thiopyrimidine derivatives as new and effective corrosion inhibitors for mild steel in hydrochloric acid: Electrochemical and quantum chemical studies. *Journal of the Taiwan Institute of Chemical Engineers*, 60, 588-601.
- [54] Anand, A., Patience, A. A., Sharma, N., & Khurana, N. (2017). The present and future of pharmacotherapy of Alzheimer's disease: A comprehensive review. *European journal of pharmacology*, 815, 364-375.
- [55] Saxena, A., Prasad, D., Haldhar, R., Singh, G., & Kumar, A. (2018). Use of Sida cordifolia extract as green corrosion inhibitor for mild steel in 0.5 M H2SO4. *Journal of environmental chemical engineering*, 6(1), 694-700.
- [56] Ahmadi, M. H., Ghazvini, M., Sadeghzadeh, M., Alhuyi Nazari, M., Kumar, R., Naeimi, A., & Ming, T. (2018). Solar power technology for electricity generation: A critical review. *Energy Science & Engineering*, 6(5), 340-361.
- [57] Kant, N., Wani, M. A., & Kumar, A. (2012). Self-focusing of Hermite–Gaussian laser beams in plasma under plasma density ramp. *Optics Communications*, 285(21-22), 4483-4487.
- [58] Gupta, V. K., Sethi, B., Upadhyay, N., Kumar, S., Singh, R., & Singh, L. P. (2011). Iron (III) selective electrode based on S-methyl N-(methylcarbamoyloxy) thioacetimidate as a sensing material. *Int. J. Electrochem. Sci*, 6, 650-663.
- [59] Mehta, C. M., Srivastava, R., Arora, S., & Sharma, A. K. (2016). Impact assessment of silver nanoparticles on plant growth and soil bacterial diversity. *3 Biotech*, 6(2), 254.
- [60] Gupta, V. K., Guo, C., Canever, M., Yim, H. R., Sraw, G. K., & Liu, M. (2014). Institutional environment for entrepreneurship in rapidly emerging major economies: the case of Brazil, China, India, and Korea. *International Entrepreneurship and Management Journal*, 10(2), 367-384.
- [61] Singh, A., Lin, Y., Obot, I. B., Ebenso, E. E., Ansari, K. R., & Quraishi, M. A. (2015). Corrosion mitigation of J55 steel in 3.5% NaCl solution by a macrocyclic inhibitor. *Applied Surface Science*, 356, 341-347.
- [62] Ansari, K. R., Quraishi, M. A., Singh, A., Ramkumar, S., & Obote, I. B. (2016). Corrosion inhibition of N80 steel in 15% HCl by pyrazolone derivatives: electrochemical, surface and quantum chemical studies. *RSC advances*, 6(29), 24130-24141.
- [63] Jnawali, P., Kumar, V., & Tanwar, B. (2016). Celiac disease: Overview and considerations for development of gluten-free foods. *Food Science and Human Wellness*, 5(4), 169-176.
- [64] Saggu, S., Sakeran, M. I., Zidan, N., Tousson, E., Mohan, A., & Rehman, H. (2014). Ameliorating effect of chicory (Chichorium intybus L.) fruit extract against 4-tert-octylphenol induced liver injury and oxidative stress in male rats. *Food and chemical toxicology*, 72, 138-146.

- [65] Bhatia, A., Singh, B., Raza, K., Wadhwa, S., & Katare, O. P. (2013). Tamoxifen-loaded lecithin organogel (LO) for topical application: development, optimization and characterization. *International Journal of Pharmaceutics*, 444(1-2), 47-59.
- [66] Singh, A., Lin, Y., Liu, W., Yu, S., Pan, J., Ren, C., & Kuanhai, D. (2014). Plant derived cationic dye as an effective corrosion inhibitor for 7075 aluminum alloy in 3.5% NaCl solution. *Journal of Industrial and Engineering Chemistry*, 20(6), 4276-4285.
- [67] Raza, K., Thotakura, N., Kumar, P., Joshi, M., Bhushan, S., Bhatia, A., ... & Katare, O. P. (2015). C60-fullerenes for delivery of docetaxel to breast cancer cells: a promising approach for enhanced efficacy and better pharmacokinetic profile. *International journal of pharmaceutics*, 495(1), 551-559.
- [68] Prabhakar, P. K., Prasad, R., Ali, S., & Doble, M. (2013). Synergistic interaction of ferulic acid with commercial hypoglycemic drugs in streptozotocin induced diabetic rats. *Phytomedicine*, 20(6), 488-494.
- [69] Chaudhary, A., & Singh, S. S. (2012, September). Lung cancer detection on CT images by using image processing. In 2012 International Conference on Computing Sciences (pp. 142-146). IEEE.
- [70] Mishra, V., Bansal, K. K., Verma, A., Yadav, N., Thakur, S., Sudhakar, K., & Rosenholm, J. M. (2018). Solid lipid nanoparticles: Emerging colloidal nano drug delivery systems. *Pharmaceutics*, 10(4), 191.
- [71] Singh, A. (2012). Hydroxyapatite, a biomaterial: its chemical synthesis, characterization and study of biocompatibility prepared from shell of garden snail, Helix aspersa. *Bulletin of Materials Science*, *35*(6), 1031-1038.
- [72] Arora, S., & Anand, P. (2019). Binary butterfly optimization approaches for feature selection. *Expert Systems with Applications*, 116, 147-160.
- [73] Chhikara, N., Kushwaha, K., Sharma, P., Gat, Y., & Panghal, A. (2019). Bioactive compounds of beetroot and utilization in food processing industry: A critical review. *Food Chemistry*, 272, 192-200.
- [74] Singh, S., Kumar, V., Chauhan, A., Datta, S., Wani, A. B., Singh, N., & Singh, J. (2018). Toxicity, degradation and analysis of the herbicide atrazine. *Environmental chemistry letters*, 16(1), 211-237.
- [75] Baranwal, T., & Pateriya, P. K. (2016, January). Development of IoT based smart security and monitoring devices for agriculture. In 2016 6th International Conference-Cloud System and Big Data Engineering (Confluence) (pp. 597-602). IEEE.
- [76] Trukhanov, S. V., Trukhanov, A. V., Salem, M. M., Trukhanova, E. L., Panina, L. V., Kostishyn, V. G., ... & Sivakov, V. (2018). Preparation and investigation of structure, magnetic and dielectric properties of (BaFe11. 9Al0. 1019) 1-x-(BaTiO3) x bicomponent ceramics. Ceramics International, 44(17), 21295-21302.
- [77] Singh, S., Singh, N., Kumar, V., Datta, S., Wani, A. B., Singh, D., ... & Singh, J. (2016). Toxicity, monitoring and biodegradation of the fungicide carbendazim. *Environmental chemistry letters*, 14(3), 317-329.
- [78] Bhyan, B., Jangra, S., Kaur, M., & Singh, H. (2011). Orally fast dissolving films: innovations in formulation and technology. *Int J Pharm Sci Rev Res*, 9(2), 9-15.
- [79] Saxena, A., Prasad, D., Haldhar, R., Singh, G., & Kumar, A. (2018). Use of Saraca ashoka extract as green corrosion inhibitor for mild steel in 0.5 M H2SO4. *Journal of Molecular Liquids*, 258, 89-97.
- [80] Panghal, A., Janghu, S., Virkar, K., Gat, Y., Kumar, V., & Chhikara, N. (2018). Potential non-dairy probiotic products—A healthy approach. *Food bioscience*, 21, 80-89.

- [81] Kumar, D., Agarwal, G., Tripathi, B., Vyas, D., & Kulshrestha, V. (2009). Characterization of PbS nanoparticles synthesized by chemical bath deposition. *Journal of Alloys and Compounds*, 484(1-2), 463-466.
- [82] Ansari, K. R., Quraishi, M. A., & Singh, A. (2015). Corrosion inhibition of mild steel in hydrochloric acid by some pyridine derivatives: an experimental and quantum chemical study. *Journal of Industrial and Engineering Chemistry*, 25, 89-98.
- [83] Singh, P. S., Singh, T., & Kaur, P. (2008). Variation of energy absorption buildup factors with incident photon energy and penetration depth for some commonly used solvents. *Annals of Nuclear Energy*, 35(6), 1093-1097.
- [84] Ansari, K. R., Quraishi, M. A., & Singh, A. (2015). Isatin derivatives as a non-toxic corrosion inhibitor for mild steel in 20% H2SO4. *Corrosion Science*, 95, 62-70.
- [85] Singh, A., Lin, Y., Ebenso, E. E., Liu, W., Pan, J., & Huang, B. (2015). Gingko biloba fruit extract as an eco-friendly corrosion inhibitor for J55 steel in CO2 saturated 3.5% NaCl solution. *Journal of Industrial and Engineering Chemistry*, 24, 219-228.
- [86] Dey, A., Bhattacharya, R., Mukherjee, A., & Pandey, D. K. (2017). Natural products against Alzheimer's disease: Pharmaco-therapeutics and biotechnological interventions. *Biotechnology Advances*, 35(2), 178-216.
- [87] Ansari, K. R., Quraishi, M. A., & Singh, A. (2015). Pyridine derivatives as corrosion inhibitors for N80 steel in 15% HCl: Electrochemical, surface and quantum chemical studies. *Measurement*, 76, 136-147.
- [88] Patel, S. (2012). Threats, management and envisaged utilizations of aquatic weed Eichhornia crassipes: an overview. Reviews in Environmental Science and Bio/Technology, 11(3), 249-259.
- [89] Mia, M., Gupta, M. K., Singh, G., Królczyk, G., & Pimenov, D. Y. (2018). An approach to cleaner production for machining hardened steel using different cooling-lubrication conditions. Journal of Cleaner Production, 187, 1069-1081.
- [90] Kondrateva T.S. Biopharmaceutical studies of children's suppositories with phosphothiamine. Pharmacy.-Moscow, 1990.-No.5.-P.14-15.
- [91] Maksudova F.Kh., Karieva E.S., Tursunova M.Kh. Study of the pharmacological properties of the combined gel of sodium diclofenac and benzketozone./Infection, immunity and pharmacologists I.- Tashkent.-2015.-No.C.160-163/
- [92] Maksudova F. Kh., Karieva E. S. In vitro equivalence evaluationce of diclofenac sodium generic medicinal preparation. // Pharmacy, a scientific and practical journal, special issue, St. Petersburg, 2016, pp. 461-464.
- [93] Piotrovsky V.K. Model and model-independent methods for describing pharmacokinetics: advantages, disadvantages and interrelation. // Antibiotics and medical biotechnology. -Moscow, 1997.-№7.P.492-497.
- [94] Kukes V.G., Sychev D.A. Clinical pharmacology. 5th ed., Moscow, 2017, p. 478.
- [95] Tillaeva U. M., Azizov U. M. Development of a methodology for isolating the amount of fensulcal determination from a biological object. Materials of the scientific-practical conference "Actual issues of education, science and production in pharmacy. Tashkent, 2009.-P.172.
- [96] Tillaeva U.M. Standardization and quality control of fensulcal in soft dosage forms. // Authors' dissertation for the study of the academician of the candidate of pharmaceuticals. Sciences . Tashkent. 2011.23 s.
- [97] Golovkin V.A. On the importance of pharmacokinetics modeling for increasing the efficiency of biopharmaceutical research. // Optimization of drug supply and ways to increase the effectiveness of pharmaceutical science: Sat. Tez.dokl.-Kharkov, 1986.-P.61-62.
- [98] Stefanova A.V. Preclinical studies of medicines. Kiev. -2002. -650 p.

- [99] L.V. Adamyan M Yoma uterus: diagnosis, treatment and rehabilitation. Clinical practice guidelines for the management of patients (draft). Moscow, 2015. Pp. 8-11.
- [100] Badmaeva S.Zh., Tskhai V.B., Grigoryan E.S., Polstyanaya G.N. M Yoma uterus: modern aspects of the etiology and pathogenesis (review). Mother and child in Kuzbass. No. 1 (76) 2019. Pages. 4-9.
- [101] Buyanova SN, Shchukina NA, Babunashvili EL .. Biological role, therapeutic potential of phytoflavonoids, vitamin D in the treatment of patients with uterine fibroids and other promising pharmacological directions. 2018 pp. 71-86. https://orcid.org/0000-0002-1358-6640.
- [102] Irnazarova D.Kh. Vitamin D supply for women with uterine fibroids. Medical journal. ISSN 0025-830 the X . 2020, no. 1, pp. 45-54.
- [103] Povoroznyuk V.V., Pludovski P., Balatskaya N.I. and other Vitamin D deficiency and deficiency: epidemiology, diagnosis, prevention and treatment. Kiev, 2015 .-- 262 p.
- [104] Radzinsky V.E., Arkhipova M.P. Uterine myoma: Problems and perspectives of the beginning of the century Medical Council, 2014. Pp. 30-32.
- [105] Yuldasheva D.Yu., Kayumova D.T., Irnazarova D.Kh., Kamilova I.A. Treatment algorithm for uterine fibroids. M etodicheskie recommendations. Tashkent, 2020. Page 30.
- [106] Ayman Al-Hendy, Marwa Badr. Can vitamin D reduce the risk of uterine fibroids? Women's Health July (2014) 10 (4), 353–358. DOI: 10.2217 / whe.14.24.
- [107] Baird DD, Hill MC, Schectman JM, et al. Vitamin D and the risk of uterine fibroids. Epidemiology, 2014; 24: 447-53. doi: 10.1097 / EDE.0b013e31828acca0.
- [108] Blauer M., Rovio PH, Ylikomi T., Heinonen PK Vitamin D inhibits myometrial and leiomyoma cell proliferation in vitro. Fertil Steril 2009; 91: 1919-25.
- [109] Brakta S., Diamond JS, Al-Hendy A., et al. Role of vitamin D in uterine fibroid biology. Fertil. Steril. 2015; 104 (3): 698-706.
- [110] Dora Pavone, Sara Clemenza, Flavia Sorbi, Massimiliano Fambrini, Felice Petraglia. Epidemiology and Risk Factors of Uterine Fibroids. Best Practice & Research Clinical Obstetrics and Gynaecology 46 (2018) 3-11. https://doi.org/10.1016/j.bpobgyn.2017.09.004.
- [111] Geum Seon Sohn, SiHyun Cho, Yong Man Kim, Chi-Heum Cho, Mee-Ran Kim, Sa Ra Lee. Current medical treatment of uterine fibroids. Obstetrics and Gynecology Science 2018; 61 (2): 192-201. https://doi.org/10.5468/ogs.2018.61.2.192.
- [112] Hajhashemi M., Ansari M., Haghollahi F., Eslami B. The effect of vitamin D supplementation on the size of uterine leiomyoma in women with vitamin D deficiency. Caspian J Intern Med 2019; 10 (2): 125-131. Doi: 10.22088 / cjim.10.2.125.
- [113] Irnazarova DH, Yuldasheva D.Yu., Najmutdinova DK, Kayumova DT, Sadikova DR, Atahodjayeva FA, Ahmedova G. Vitamin D status in women with uterine fibroids (uf) of the uzbek population. JOURNAL OF CRITICAL REVIEWS. ISSN- 2394-5125 VOL 7, ISSUE 17, June 2020. p. 2240-2250. http://www.jcreview.com/?mno=96009
- [114] Irnazarova D.Kh. Innovative technologies in the diagnostics of uterine fibroids. R e-Health Journal Scientific and practical journal. ISSN 2181-0443. Andijan 2020, issue: 4. Pages. 29.
- [115] John L. Wu, James H. Segars. Is vitamin D the answer for prevention of uterine fibroids? Fertility and Sterility, Vol. 104 No. 3 / September 2015. P. 559-560. http://dx.doi.org/10.1016/j.fertnstert.2015.06.034.
- [116] Maria Syl D. De La Cruz, Edward M. Buchanan. Uterine Fibroids: Diagnosis and Treatment. American Family Physician . 2017; 95 (2): 100-107.

- [117] Mohamed Ali, Ayman Al-Hendy, Qiwei Yang. Vitamin D, a promising natural compound with anti-uterine fibroid characteristics. Fertility and Sterility, Vol. 111 N. 2 / February 2019. Https://doi.org/10.1016/j.fertnstert.2018.11.004.
- [118] Mohanambal M. Munusamy, Wills G. Sheelaa, Vijaya P. lakshmi. Clinical presentation and prevalence of uterine fibroids: a 3-year study in 3-decade rural South Indian women. Int J Reprod Contracept Obstet Gynecol. 2017 Dec; 6 (12): 5596-5601. DOI: http://dx.doi.org/10.18203/2320-1770.ijrcog20175288]
- [119] Paffoni A., Somigliana E., Vigano P., et al. Vitamin D status in women with uterine leiomyomas. J Clin Endocrinol. Metab. 2013; 98 (8): E1374-8.
- [120] Ren Jinhe. Color Doppler Ultrasound in Uterine Arterial Embolization. Open Med. 2017; 12: 489-493. https://doi.org/10.1515/med-2017-0069.
- [121] Sabry M., Halder SK, Allah AS, et al. Serum vitamin D3 level inversely correlates with uterine fibroid volume in different ethnic groups: a cross-sectional observational study. Int J Womens Health 2013; 5: 93-100.
- [122] Scientific Advisory Committee on Nutrition. Vitamin D and Health, 2016. P 11-12. https://www.gov.uk/government/groups/scientific-advisory-committee-on-nutrition.
- [123] Xess S., Sahu J. International Journal of Reproduction, Contraception, Obstetrics and Gynecology . 2020 Apr; 9 (4): 1477-1481. DOI: http://dx.doi.org/10.18203/2320-1770.ijrcog20201208 .