ATTN: THE CHAIRMAN OF THE SCIENTIFIC JURY AT THE MEDICAL UNIVERSITY "PROFESSOR PARASKEV STOYANOV" OF VARNA

Concerning:

the participation of Assoc. Prof. Miglena Dimitrova Georgieva, MD, PhD, in the competition promulgated in *State gazette* No 61/July 23, 2021 for tenure of the academic position of "Professor" in the field of higher education No 7 Public health and sports, professional trend No 7.1 Medicine and scientific speciality of Pediatrics, at 0,5 pay-roll office for the needs of the Department of Pediatrics at the Faculty of Medicine of the Medical University "Prof. Paraskev Stoyanov" of Varna and 0,5 pay-roll office for the needs of the Second Clinic of Pediatrics at St. Marina University Hospital of Varna, according to Order No R-109-422/September 27, 2021 of the Rector of the Medical University "Prof. Paraskev Stoyanov" of Varna, Prof. Valentin Lyubomirov Ignatov, MD, PhD.

PEER REVIEW

by

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According to Protocol No 1/September 30, 2021 of the Scientific jury, appointed according to Order of the Rector of the Medical University "Prof. Paraskev Stoyanov" of Varna Prof. Valentin Lyubomirov Ignatov, MD, PhD, I have been assigned to prepare the present peer review for the only candidate in the competition.

1. CONCISE BIOGRAPHIC AND PROFESSIONAL DATA

Assoc. Prof. Miglena Dimitrova Georgieva, MD, PhD, was born on March 14, 1961in the city of Varna. She graduated at Frédéric Joliot-Curie Fourth Foreign language secondary school with excellent marks and gold medal in 1980 and at Leningrad Pediatric Medical Institute with excellent marks in 1986. In 1987, she started to work as intern in the Pediatric Division of the Regional Hospital of the town of Provadiya. During the period between 1992 and 1996, she was regular post-graduate student in the Department of Pediatrics of the Medical University of Varna and the Clinic of Gastroenterology at St. Marina University Hospital of Varna. During the period between 1997 and 2008, she worked as intern in the Clinic of Gastroenterology and Endocrinology at the same hospital and during the period between 2008 and 2010 she worked in the Clinic of Cardiology, Rheumatology, Nephrology, Gastroenterology and Pediatric Intensive Division. Between 2010 and April 1, 2014, she worked as associate professor in the Third Division of the Multiprofile Specialized Pediatric Clinic at St. Marina University Hospital of Varna, and afterwards until presence she works as head of the Second Clinic of Pediatrics at St. Marina University Hospital of Varna. During the period between 2002 and 2015, she was part-time lecturer in the Centre for Continuing

Education, qualification of "Management in recreation" at the University of Economics of Varna. Between August 1, 2018 and March 1, 2020, she is organizer and deputy-director of the Expert Centre of Cystic Fibrosis and since March 1, 2020, she is director of the Expert Centre of Cystic Fibrosis.

Miglena Dimitrova Georgieva, MD, acquired the PhD educational and scientific degree in 1997. The theme of her dissertation work was "Clinical, endoscopic, morphological and functional alterations in upper alimentary tract diseases among children and adolescents".

She attained the academic rank of associate professor in 2010.

Specialities acquired

Assoc. Prof. Miglena Dimitrova Georgieva, MD, PhD, has a recognized speciality of pediatrics in 1992, of pediatric gastroenterology in 2006 and of nutrition and dietetics in 2018.

Oualification acquired in:

Diagnostic and therapeutic fibrogastroscopy and fibrocolonoscopy in 2008

Conventional echography (abdominal echography and superficial structures) - level one in 2008

Interventional gastrointestinal endoscopy - level two in 2016

Endoscopic cholangiopancreatography and related interventional procedures - level three in 2016

Abdominal Doppler echography in gastroenterology - level two in 2016

Interventional and endoscopic abdominal echography in gastroenterology - level three in 2016.

Postgraduate qualification

Two five-day courses of pediatric gastroenterology (chronic hepatitis and food allergy) in the city of Sofia; a ten-day course of pediatric gastroenterology (selected chapters of the pediatric gastroenterology) in the city of Krakow, Poland, within a HOPE Project; a five-day course of gastroenterology (selected chapters of the pediatric gastroenterology) within the medical seminars in the city of Salzburg, Austria; a five-day European summer school of pediatric gastroenterology (selected chapters of the pediatric gastroenterology) in the city of Sofia; a three-day European summer school of pediatric gastroenterology (selected chapters of the pediatric gastroenterology) in the city of Varna; a seven-day basic course of clinical nutrition (enteral and parenteral nutrition) in the city of Dubrovnik, Croatia; a threeday training in inflammatory bowel diseases in the city of Barcelona, Spain; a course of basic training of Gauche's disease in the city of Amsterdam, the Netherlands; five annual module trainings in nutrition and dietetics at the National Centre for Public Health Protection and Analyses in the city of Sofia; a two-month practical training in clinical nutrition and dietetics at Tsaritsa Giovanna Hospital of Sofia; a three-day training in support of the children with cystic fibrosis in the city of Southampton, Great Britain; a seven-day course of pediatric gastroenterology in the city of Graz, Austria; a course of pedagogical bases of academic teaching in the city of Varna, etc.

II. EVALUATION OF CANDIDATE'S RESEARCH ACTIVITY

1. Scientometric indicators

During the period between 1993 and 2009, Miglena Dimitrova Georgieva, MD, has a total of 173 scientific works. It deals with one dissertation for the acquisition of the educational and scientific degree of 'doctor', monograph, nine chapters in monographs and reference books, 33 articles in 10 Bulgarian journals and one article in one foreign journal, journals; 31 abstracts of reports and scientific communications published in proceedings of a total of 20 scientific forums in Bulgaria and 17 abstracts published in four in Bulgarian journals as well as 63 abstracts of reports and scientific communications published in

proceedings of a total of 38 scientific forums abroad and 17 abstracts published in seven eminent foreign journals.

The total number of the concrete scientific works of Assoc. Prof. Miglena Dimitrova Georgieva, MD, PhD, during the period between 1993 and 2921 is the following: two monographs, nine chapters in monographs and reference books, 12 publications in six proceedings of scientific forums in Bulgaria, 121 articles in 21 Bulgarian and seven foreign journals, 168 abstracts of reports and scientific communications published in proceedings of a total of 35 scientific forums in Bulgaria and of 41 ones abroad and in nine foreign journals, respectively, as well as a total of 107 plenary reports oral communications and multimedia oral presentations delivered at a total of 58 specialized scientific forums in Bulgaria and in four one abroad.

Assoc. Prof. Miglena Dimitrova Georgieva, MD, PhD, participates in the present competition with the following scientific works during the period between 2011 and 2021 after her attaining the academic rank of associate professor:

One monograph and 12 publications in 6 scientific conference proceedings in Bulgaria with a total volume of 441 pages.

A total 87 full-text articles in a total of 17 Bulgarian and 6 foreign journals with a total volume of 443 pages as the only or the first author in 38 articles (in 43,68% of the cases).

A total 40 abstracts of reports and scientific communications published in a total of 18 proceedings with abstracts (15 in Bulgaria and three abroad) and in two foreign periodical journals as the only or the first author in 38 abstracts (in 30,00% of the cases).

A total of 107 plenary reports, oral communications and multimedia oral presentations with a total volume of more than 1000 slides delivered at a total of 58 specialized scientific forums such as national and international congresses, conferences and symposia in Bulgaria and at four international scientific forum abroad (in Prague, Vienna, Jerusalem and Berlin) as the only or the first author in 59 delivered reports and scientific communications (in 55,14% of the cases).

The annual dynamics of the publication output and research activity of Assoc. Prof. Miglena Dimitrova Georgieva, MD, PhD, after her attaining the academic rank of associate professor is presented in Table 1.

Table 1. Number of published articles and abstracts as well as of the participations in scientific forums according to years

Year	articles	abstracts	participations
2011	1	-	5
2012	-	2	55
2013	9	1	10
2014	10	4	19
2015	15	15	3
2016	9	1	11
2017	9	2	9
2018	10	5	5
2019	9	5	18
2020	9	5	7
2021	6	-	15
total	87	40	107

The total impact factor after her attaining the academic rank of associate professor is 3,874 (two full-text articles).

The total number of the citations or peer reviews after her attaining the academic rank of associate professor is 54, of which 40 in scientific publications abstracted and indexed in world eminent data-bases with scientific information, or in monographs and books; two citations in scientifically peer-reviewed monographs and books and 12 citations or peer reviews in scientifically peer-reviewed journals which are not abstracted in data-bases. Three articles in three foreign journals are cited a total of 34 times while three articles in three Bulgarian journals - a total of 20 times.

2. Participations in research projects and clinical trials

Adviser of research projects in the Medical University "Prof. Paraskev Stoyanov" of Varna:

1. Assessment of the expression levels of selected microRNAs as potential biomarker in children and adolescents with inflammatory bowel diseases.

2. Bone health in children with celiakia.

3. Etiology, clinical picture and diagnosis of non-alcoholic liver disease in children and adolescents.

Participant in a research project in the Medical University "Prof. Paraskev Stoyanov" of Varna:

1. Nutritive status and food intervention in children with cystic fibrosis - a singlecentre, open non-randomized trial Participant in clinical trials

1. Improving celiac disease management in the Danube region by raising the awareness, improving the knowledge, and developing better skills.

- 2. Efficacy of Lactobacillus reuteri probiotic for prevention of antibiotic-related diarrheas and Clostridium difficile-related infections among hospitalized children and adolescents.
- 3. A phase III-B, randomized, open trial of monotherapy with pegylated interferon, alpha-2A or in combination with Lamivudine as compared to non-treated control patients - children with HBeAg-positive chronic hepatitis in a immunotolerance phase.

4. Effect of different carob mucilage contents in Friso Comfort on the lower esophageal sphincter.

5. A phase II, multicenter, parallel group, active-controlled, randomized, double-blind doseranging study to evaluate the efficacy and safety of different doses of Creon IR in subjects with pancreatic exocrine insufficiency due to cystic fibrosis.

6. A multicenter, randomized, double-blind, placebo-controlled phase II trial to evaluate the efficacy and safety of Lenbasum in cystic fibrosis.

7. A phase III, multicenter, randomized, double-blind, placebo-controlled, parallel-group efficacy and safety study of Linaclotide administered orally to children aged 6 to 17 years with functional constipation.

III. MAIN SCIENTIFIC TRENDS

The research activity of Assoc. Prof. Miglena Dimitrova Georgieva, MD, PhD, is product of an intensive clinical diagnostic and therapeutic activity as pediatrician and pediatric gastroenterologist. She has undoubted prestige not only among the colleagues in our country and abroad but also among the children with gastrointestinal diseases and functional disorders, with severe diseases of the liver, pancreas, cystic fibrosis, nutritional disorders, etc. The object of her profound investigations are not only a series of socially significant diseases but also some more seldom diseases of casuistic value for the medical science and practice. The spectrum of her scientific works is very broad. It includes populational-diagnostic and screening investigations, organization-methodological and thematic analyses as well as direct applied research in the field of pediatrics and pediatric gastroenterology.

A great number of publications in pediatric gastroenterology are devoted to actual problems in the following thematic trends: functional gastrointestinal disorders, upper alimentary tract diseases, celiakia, inflammatory and several other bowel diseases diseases as well as genetic methods for diagnosis.

In the field of pediatric hepatology and pancreatology, several chronic liver diseases, cystic fibrosis as well as acute and chronic pancreatitis are taken in consideration.

I would like to outline Assoc. Prof. Georgieva's publications on the gut microbiome, probiotics, prebiotics, psychobiotics and food allergy. In a total of 19 articles and three abstracts, the issues of the nutrition of the healthy children and of those with different diseases and functional gastrointestinal disorders are investigated. The concrete recommendations for a special nutrition in inflammatory bowel diseases, neurodegenerative diseases, cystic fibrosis, acute and chronic pancreatitis deserve attention.

In a total of 11 articles and seven abstracts, several actual issues of a series of chronic liver diseases in childhood (chronic viral and medicamentous hepatitis, Wilson's disease, α 1-antitrypsin deficiency, non-alcoholic liver disease, Gilbert's syndrome, Rey's syndrome, autoimmune liver disease, ascites syndrome, hepatorenal syndrome, acute liver failure, benign and malignant liver tumours, familial nodular hyperplasia, etc.) are considered. In five articles and one abstract, the novelties in the etiology, pathogenesis, classifications, clinical variations, diagnosis and therapeutic influence on the large bowel polyps, rotavirus and enterovirus enteritis, *Clostridium difficile* infection and diarrhea are reflected.

Two abstracts one of which has been published in an eminent foreign journal with impact factor are devoted to the acute and chronic hepatitis in children. The role of nutrition and contemporary genetic therapy of cystic fibrosis as well as the clinical peculiarities of this disease are the object of four articles and five abstracts.

In seven articles and four abstracts, the essential results from the examination of some other diseases in children such as hereditary hemorrhagic teleangiectasia, anti-NMDR encephalitis, primary form of tularemia, aspiration of foreign bodies, urticaria, headache in children with epilepsy, etc. are published.

IV. MAIN ORIGINAL SCIENTIFIC CONTRIBUTIONS

The main scientific contributions of Assoc. Prof. Miglena Georgieva, MD, PhD, are in the field of the diagnosis, clinical picture and treatment of the functional gastrointestinal disorders, celiakia, cystic fibrosis, inflammatory and other bowel diseases, chronic liver disease (viral hepatitis and metabolic liver diseases, especially, non-alcoholic fatty liver disease), probiotics, prebiotics and psychobiotics, nutrition of the healthy suckling and child as well as in case of several diseases such as celiakia, cystic fibrosis, inflammatory bowel diseases, disorders in the autistic spectrum, neurodegenerative diseases and food allergy, rare diseases of the alimentary system and other organs and the genetic methods for the proof of the gastrointestinal tract diseases as well.

In general, the theoretical and methodological and practically applied contributions in the scientific works of Assoc. Prof. Miglena Georgieva, MD, PhD, could be systematized in the following main thematic fields: pediatric gastroenterology, pediatric hepatology and pancreatology, nutrition, clinical pediatrics, and casuistry.

The problem of the functional gastrointestinal disorders is comprehensively considered for the first time in Bulgaria in an independent monograph and six articles. The classification of Rome IV criteria is introduced and algorithms for the diagnosis and schedules for non-medicamentous and drug treatment of these patients.

The diagnosis of the gastroesophageal reflux by means of impedance pH-metry is used for the first time in Bulgaria in two articles and one abstract. Both one article and one abstract are devoted to the clinical picture, diagnosis and treatment of the achalasia.

For the first time in Bulgaria, genetic tests for the proof of the genetic predisposition towards celiakia are applied. Seven articles and six abstracts are devoted to the diagnosis and treatment of this disease.

The microribonucleic acids are used for the first time in Bulgaria in the diagnosis and follow-up of two inflammatory bowel diseases such as ulcerative colitis and Crohn's disease in childhood and the results are published in three articles. In other eight articles and three abstracts, the issues of the deep venous thrombosis and the recommendations of the European Society of Pediatric Gastroenterology, Hepatology and Nutrition concerning the role of the nutrition in the treatment of these diseases in childhood are discussed.

In seven articles and one abstract, the contemporary concepts of the intestinal microflora, pro-, pre- syn- and psychobiotics for the first time in Bulgaria are systematized and the positive prophylactic and therapeutic effect of the probiotics in the diarrhoea related with antibiotic therapy is proved.

In four articles and one abstract for the first time in Bulgaria, the role of the extensively hydrolyzed milks for the treatment of suckling allergy towards cow milk as well as of the partially hydrolyzed milks for food allergy prevention is described. In four articles and four abstracts for the first time in Bulgaria, the results from genetic investigations of the HLA-genotypification in celiakia and suspicion on Gilbert's disease are published. For the first time in Bulgaria, the results about adipokine and other markers of metabolic syndrome and abdominal obesity are published.

It could be generalized that up Assoc. Prof. Miglena Georgieva, MD, PhD, succeeds not only in implementing the timely and precise diagnosis of the mentioned diseases and disorders in sucklings, children and adolescents but also in performing the most appropriate individualized treatment that conforms on the standards for best clinical practice. Her efforts are directed not only to relief ill child's suffering but also to avoid the severe socially significant diseases at later age. She popularizes her accumulated clinical experience by mens of a scientific production of impressing volume and high quality.

V. TEACHING, LECTURER'S AND PUBLIC ACTIVITY

The teaching loading of Assoc. Prof. Miglena Georgieva, MD, PhD, during the last four academic years is the following: during the 2017-2018 academic year - 144 hours, during the 2018-2019 academic year - 136 hours, 144 hours, and during the 2019-2020 academic year - 124 hours.

She is adviser of three postgraduate students in pediatrics and of four postgraduate students in pediatric gastroenterology.

Assoc. Prof. Miglena Georgieva, MD, PhD, performs education according to individual courses for postgraduate qualification in pediatric gastroenterology and nutrition.

Under the guidance of Assoc. Prof. Miglena Georgieva, MD, PhD, two postgraduate students, Niya Yolkova Rasheva, MD, and Krasimira Ivanova Koleva, MD, have defended their doctoral theses for the acquisition of the educational and scientific degree of PhD. Nowadays, she is adviser of six other postgraduate students - Yoana Dyankova Dyankova, MD, Margarita Ivanova Nikolova, MD, Mihail Fotinov Siromahov, MD, Zhivka Emilova Chuperkova, MD, Martina Ivanova-Galabova, MD and Diana Yordanova Yordanova, MD.

Assoc. Prof. Miglena Dimitrova Georgieva, MD, PhD, is member of the Board of managers of the Bulgarian Pediatric Association and president of the Bulgarian Society of Pediatric Gastroenterology, Hepatology and Nutrition. She is member of the following professional and scientific organizations:

- 1. Bulgarian Pediatric Association
- 2. Bulgarian Scientific Society of Gastroenterology
- 3. Bulgarian Union of Physicians

- 4. BULSPEN (Bulgarian Society of Enteral and Parenteral Nutrition)
- 5. Bulgarian Society of Neurogastroenterology and Motility
- 6. ICMART (International Council of Medical Acupuncture and Related Techniques)
- 7. Société Française de Pédiatrie
- 8. ESPGHAN (European Society of Pediatric Gastroenterology, Hepatology and Nutrition)
- 9. ESPEN (European Society of Enteral and Parenteral Nutrition).

 She has participated in the scientific jury for awarding of the educational and scientific degree of PhD by six postgraduate students, for two candidates for "associate professor" and "professor" each and in the election of one "assistant-in-chief".

CONCLUSION

Based on the aforementioned facts, I can declare that the professional qualification, research as well as teaching and lecturer's activity of Assoc. Prof. Miglena Dimitrova Georgieva, MD, PhD, entirely meet the requirements of the Statute-book for the conditions and order for the acquisition of scientific degrees and for tenure of the academic position of "professor" in the Medical University "Prof. Paraskev Stoyanov" of Varna. That is why I recommend to the honoured members of the Scientific jury to vote positively for the tenure by Assoc. Prof. Miglena Dimitrova Georgieva, MD, PhD, of the academic position of "Professor" in the Medical University "Prof. Paraskev Stoyanov" of Varna for the needs of the Department of Pediatrics at the Faculty of Medicine of the Medical University Prof. Paraskev Stoyanov of Varna and the Second Clinic of Pediatrics at St. Marina University Hospital of Varna.

Varna, October 18, 2021

Peer reviewer: (Prof. Nikola Kolev, MI), Phi DSc)