

CLARE L. FRASER^f
 RUTH HUNA-BARON^g
 BRADLEY KATZ^e
 MITCHELL LAWLOR^h
 LASSE MALMQVISTⁱ
 NANCY J. NEWMAN^b
 JASON PERAGALLO^a
 AXEL PETZOLD^j
 PREM S. SUBRAMANIAN^k
 JUDITH E.A. WARNER^e
 MARIANNE WEGENERⁱ
 SUI WONG^j
 STEFFEN HAMANNⁱ

ODDS CONSORTIUM
^aDepartment of Ophthalmology
 State University of New York at Stony Brook
 Health Sciences Center
 Stony Brook, New York, USA

^bEmory University School of Medicine
 Atlanta, USA
^cIvey Eye Institute
 Western University
 London, Ontario, Canada

^dDepartment of Clinical Neurosciences
 University of Calgary
 Calgary, Canada

^eUniversity of Utah
 Department of Ophthalmology
 Moran Eye Center
 Salt Lake City, Utah, USA

^fSave Sight Institute
 University of Sydney
 Australia

^gSheba Medical Center
 Tel Hashomer, Israel
^hSydney Eye Hospital
 University of Sydney
 Australia

ⁱDepartment of Ophthalmology
 Rigshospitalet, University of Copenhagen
 Glostrup, Denmark

^jMoorfields Eye Hospital
 London, UK

^kUniversity of Colorado School of Medicine
 Denver, Colorado, USA

REFERENCES

1. Kim MS, Lee KM, Hwang JM, et al. Morphologic features of buried optic disc drusen on en face optical coherence tomography and optical coherence tomography angiography. *Am J Ophthalmol* 2020;213:125–133.
2. Malmqvist L, Bursztyn L, Costello F, et al. The Optic Disc Drusen Studies Consortium recommendations for diagnosis of optic disc drusen using optical coherence tomography. *J Neuroophthalmol* 2018;38:299–307.

3. Lee KM, Woo SJ, Hwang JM. Morphologic characteristics of optic nerve head drusen on spectral-domain optical coherence tomography. *Am J Ophthalmol* 2013;155:1139–1147.
4. Lee KM, Woo SJ, Hwang JM. Peripapillary hyperreflective ovoid mass-like structures: is it optic disc drusen or not? [letter to editor and author response]. *J Neuroophthalmol* 2018;38:567–570.
5. Malmqvist L, Sibony PA, Fraser CL, et al. Peripapillary ovoid hyperreflectivity in optic disc edema and pseudopapilledema. *Ophthalmology* 2018;125:1662–1664.
6. Pichi F, Romano S, Villani E, et al. Spectral-domain optical coherence tomography findings in pediatric tilted disc syndrome. *Graefes Arch Clin Exp Ophthalmol* 2014;252:1661–1667.
7. Lyu In Jeong, Park Kyung-Ah, Oh Sei Yeul. Association between myopia and peripapillary hyperreflective ovoid mass-like structures in children. *Scientific Reports* 2020; 10:2238. <https://doi.org/10.1038/s41598-020-58829-3>.

Reply to Comment on: Morphologic Features of Buried Optic Disc Drusen on En Face Optical Coherence Tomography and Optical Coherence Tomography Angiography



WE GREATLY APPRECIATE THE INTEREST THAT SIBONY and associates¹ demonstrated in our study. The authors raised an important question regarding the definition of optic disc drusen (ODD) in our study. We know that the Optic Disc Drusen Studies Consortium defined ODD as hyporeflective lesions surrounded by hyperreflective capsules using optical coherence tomography (OCT). We previously showed that this kind of visible drusen can be observed especially in the eyes of the elderly associated with degenerative changes on OCT.² However, in young patients, ODD can be seen as a hyperreflective mass on OCT, located nasal to the optic disc. The Consortium denies the latter ODD as true drusen and named them peripapillary ovoid mass-like structures (PHOMS) instead. PHOMS is a vague term denying any cause for or mechanism of the lesion. The most important reason why the Consortium used the term PHOMS instead of ODD is based on the histopathological examination described in their paper in 2018 (see their Figure 4C-E).³ However, we are still doubtful about the histological images, and the true pathogenesis of PHOMS or buried ODD (by our definition) remains to be defined.⁴

To prove whether PHOMS is merely herniated optic nerve fibers or not, we analyzed the en face OCT images in this study.¹ Cross-sectional OCT, en face OCT, and OCT angiography simulate in vivo histology and could be superior to ex vivo histopathology in certain aspects, as the last could be influenced by postmortem changes

and preparation artifacts. In our study, there is no evidence suggesting that PHOMS is composed of herniated optic nerve fibers. Rather, it is a mass lesion clearly distinct from optic nerve fibers with poor vascularity, suggesting an accumulation of extracellular deposits. Therefore, we called it ODD in our study.

We speculated that the mechanism of buried ODD or PHOMS is the accumulation of intracellular microorganisms from disturbed axonal flow of the optic nerve. Sibony and associates¹ suggested 3 kinds of PHOMS in their comment. The 3-PHOMS categories perfectly explains the condition of disturbed axonal flow. Our previous study results showing that ODD occurs in eyes with small optic discs and small scleral canals also support the mechanism of buried ODD or PHOMS as the accumulation of axoplasmic materials.^{2,5}

To finalize this dispute, more histology data regarding PHOMS or buried ODD should be obtained and analyzed. Another way to prove that PHOMS is a feature of ODD is to show the de novo development of PHOMS in eyes without ODD. In addition, long-term follow-up of PHOMS or buried ODD using en face OCT could provide additional evidence of its true pathophysiology and its association with visible ODD.

Finally, we want to highlight our main study conclusion that en face OCT and OCT angiography could be used to differentiate optic disc edema from pseudopapilledema regardless of whether it is ODD or PHOMS.

MIN SEOK KIM
Seongnam, Republic of Korea
KYOUNG MIN LEE
Seoul, Republic of Korea

JEONG-MIN HWANG
HEE KYUNG YANG
SE JOON WOO
Seongnam, Republic of Korea

ALL AUTHORS HAVE COMPLETED AND SUBMITTED THE ICMJE form for Disclosure of Potential Conflicts of Interest and none were reported.

Funding/Support: No outside funds were received for this work.

Financial disclosures: S.J.W. is a consultant for Samsung Bioepis (Songdo, South Korea), Panolos Bioscience (Seoul, South Korea), and Novelty Nobility (Seoul, South Korea); and is a cofounder of RetiMark, Seoul, South Korea. All other authors have reported that they have no relationships relevant to the contents of this paper to disclose.

REFERENCES

1. Kim MS, Lee KM, Hwang JM, Yang HK, Woo SJ. Morphologic features of buried optic disc drusen on en face optical coherence tomography and optical coherence tomography angiography. *Am J Ophthalmol* 2020;213:125–133.
2. Lee KM, Woo SJ, Hwang JM. Morphologic characteristics of optic nerve head drusen on spectral-domain optical coherence tomography. *Am J Ophthalmol* 2013;155:1139–1147.
3. Malmqvist L, Bursztyn L, Costello F, et al. The optic disc drusen studies consortium recommendations for diagnosis of optic disc drusen using optical coherence tomography. *J Neuroophthalmol* 2018;38:299–307.
4. Tso MO. Pathology and pathogenesis of drusen of the optic nerve head. *Ophthalmology* 1981;88:1066–1080.
5. Lee KM, Woo SJ, Hwang JM. Differentiation of optic nerve head drusen and optic disc edema with spectral-domain optical coherence tomography. *Ophthalmology* 2011;118:971–977.