Uveitis- a rare disease often associated with systemic diseases and infections- a systematic review of 2619 patients

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Abstract

Background: Uveitis is an autoimmune disease of the eye that refers to any of a number of intraocular inflammatory conditions. Because it is a rare disease, uveitis is often overlooked, and the possible associations between uveitis and extra-ocular disease manifestations are not well known. The aim of this study was to characterize uveitis in a large sample of patients and to evaluate the relationship between uveitis and systemic diseases.

Methods: The present study is a cross-sectional study of a cohort of patients with uveitis. Records from consecutive uveitis patients who were seen by the Uveitis Service in the Department of Ophthalmology at the Medical University of Vienna between 1995 and 2009 were selected from the clinical databases. The cases were classified according to the Standardization of Uveitis Nomenclature Study Group criteria for Uveitis.

Results: Data were available for 2619 patients, of whom 59.9% suffered from anterior, 14.8% from intermediate, 18.3% from posterior and 7.0% from panuveitis. 37.2% of all cases showed an association between uveitis and extra-organ diseases; diseases with primarily arthritic manifestations were seen in 10.1% of all cases, non-infectious systemic diseases (i.e., Behçet’s disease, sarcoidosis or multiple sclerosis) in 8.4% and infectious uveitis in 18.7%. 49.4% of subjects suffering from anterior uveitis tested positively for the HLA-B27 antigen. In posterior uveitis cases 29% were caused by ocular toxoplasmosis and 17.7% by multifocal choroiditis.

Conclusion: Ophthalmologists, rheumatologists, infectiologists, neurologists and general practitioners should be familiar with the differential diagnosis of uveitis. A better interdisciplinary approach could help in tailoring of the work-up, earlier diagnosis of co-existing diseases and management of uveitis patients.

Keywords: Uveitis, Etiology, Systemic associations, Arthritis, Infections

Background

Uveitis is a sight-threatening inflammation inside the eye that affects both the uveal tract (which is composed of the iris, choroid, and ciliary body and which is the blood-supplying layer inside of the eye), and adjacent structures (including the sclera, cornea, vitreous humor, retina and optic nerve head). Because the disease involves recurrent intraocular inflammation, uveitis can cause transient or permanent visual impairment and ocular complications that are not responsive to therapy [1-4]. Uveitis can occur either as a co-manifestation of various autoimmune disorders and infections or as a side effect of medications and toxins, or it can arise as a purely idiopathic ocular inflammation [3,5-10].

The prevalence of uveitis is estimated at 38 cases per 100,000 people, so it meets the criteria for classification as a rare disease [2,11-19]. It is particularly prevalent in younger people; the mean age of uveitis patients at the onset of the disease is less than 40 years of age [20-22].
Although it is an orphan disease, uveitis is the fourth most common cause of blindness among the working-age population in the developed world, and its economical and social impact not yet been evaluated [3,20-22].

The referral of a patient to a uveitis expert is often delayed because uveitis is commonly unknown, and it is therefore under-recognized. This delay in diagnosis and referral increases the risk that uveitis will result in irreversible damage to various ocular structures. Prompted by the hypothesis that highlighting systemic associations would optimize the referral of patients with uveitis, we conducted this systematic review to analyze the distributions of uveitis subtypes and their extra-organ manifestations.

**Patients and methods**

The institutional review board of the Medical University of Vienna approved this study. Since September 1995, data from all patients referred to the Uveitis Unit of the Department of Ophthalmology, Medical University of Vienna, were systematically recorded. A total of 2619 consecutive patient records were analyzed for this study.

The uveitis subtypes were classified based on the specific disease patterns and diagnoses following the recommendations of the International Uveitis Study Group and the Standardization of Uveitis Nomenclature (SUN) working group [23,24]. Guided by the medical history of each patient, investigations of the anatomical location and character of the inflammation as well as subsequent diagnostics were performed using a tailored approach [25,26]. All patients were treated in a multidisciplinary setting and referred to the respective specialist if systemic disease was suspected.

Factors considered to satisfy the criteria of specific diagnosis included well-defined a etiology, typical clinical appearance and history or classification based on pathological or pathognomonic laboratory parameters.

All patients with acute anterior uveitis were typed selectively for presence of the HLA-B27 antigen, as HLA-B27–associated acute anterior uveitis with or without systemic disease is recognized as a specific uveitis entity [27]. All uveitis patients with multifocal choroiditis showing the typical clinical appearance of birdshot choroidopathy were typed for the HLA-A29 antigen [28]. These two HLA-antigen predispositions are considered to be pathognomonic when combined with the typical clinical appearance.

Diseases belonging to the white dot syndromes which will be discussed in this article are acute posterior multifocal placoid pigment epitheliopathy (APMPPE), multiple evanescent white dot syndrome (MEWDS), birdshot retinochoroidopathy (BSRC), multifocal choroiditis (MFC), punctuate inner choroidopathy (PIC), acute zonal occult outer retinopathy (AZOOR) and serpiginous choroiditis.

Descriptive statistics were computed using SPSS 17.0. Values are given as ranges, percentages or means ± standard error of the mean, as applicable.

**Results**

The patient sample comprised 1339 women and 1278 men (51.1% and 48.2%, respectively). Uveitis occurred at a mean age of 38.8 ± 18.3 years (range 1-87). The age at the onset of the disease was between 17-60 years in 79.8% of the patients. 9.2% of all cases were children (<17 years) and 11% were elderly patients (>60 years).

The anatomical classification of uveitis was anterior in 59.9%, intermediate in 14.8%, posterior in 18.3% and panuveitis in 7% of cases. Figure 1 summarizes the distribution of unclassified and classified cases in these four groups.

**Specific diagnoses**

Diseases with primarily arthritic manifestations and uveitis were diagnosed in 265 cases (10.1%). Of these, the specific aetiologies included enclosing spondylitis (n = 143), rheumatoid arthritis (n = 42), juvenile idiopathic arthritis (JIA) (n = 58), reactive arthritis (n = 5) and psoriatic arthritis (n = 17).

Infectious uveitis was diagnosed in 495 patients (19.0%). Diagnoses included 226 viral [ocular herpes virus reactivation

![Figure 1: Percentage of unclassified and classified cases according to anatomical localization of uveitis.](image-url)
systemic varicella virus infection (VZV) (n = 14), systemic cytomegalovirus infection (CMV) (n = 5), other (n = 10)], 198 parasitic [toxoplasmosis (n = 173), 18 toxocarosis (n = 18), other (n = 7)], 47 bacterial and 24 fungal infections. Of these cases, 99 were associated with current systemic infections (Table 1).

Of the purely ocular infections, 187 cases were diagnosed with herpetic anterior uveitis, 14 with acute retinal necrosis (ARN), 173 with ocular toxoplasmosis and 18 with ocular toxocarosis.

Non-infectious systemic diseases with primarily non-arthritic manifestations were diagnosed in 221 cases (8.4%). These included Behçet’s disease (n = 49), sarcoidosis (n = 64), Crohn’s disease (n = 30), multiple sclerosis (MS) (n = 25), ulcerative colitis (n = 14), Whipple’s disease (n = 2), Vogt-Koyanagi-Harada disease (VKH) (n = 11), systemic lupus erythematosus (SLE) (n = 7) and masquerade syndromes (MASQ) (n = 19).

Ocular syndromes such as HLA-B27–associated anterior uveitis without systemic disease (HLA-B27 AAU) (n = 271), Fuchs’ heterochronic cyclitis (HCF) (n = 88), serpiginous choroidopathy (n = 23), multifocal choroiditis (n = 86), birdshot choroidopathy (n = 10), acute posterior multifocal placoid pigment epitheliopathy (APMPPE) (n = 8), multiple evanescent white dot syndrome (n = 6), granulomatous cyclic crisis (n = 10) and sympathetic ophthallmia (n = 2), were found in 503 (19.2%) cases.

Anatomical location and specific entities

In the anterior uveitis (AU) group, specific diagnoses could be established in 60.9% of cases. HLA-B27–associated anterior uveitis was diagnosed in 414 cases (30.5% of AU). These cases were either related to underlying HLA-B27–associated systemic diseases such as enclosing spondylitis (n = 143) or to purely ocular disease (271 cases).

Herpetic anterior uveitis and HCF were found in 11.4% and 4.5% of cases, respectively. JIA was seen in 3.4% of AU patients. In addition, uveitis was associated with sarcoidosis in 2.1% of cases, and Behçet’s disease in 0.9%. Systemic infections, MASQ, granulomatous cyclic crisis, and ulcerative colitis were each seen in less than 1% of AU patients.

75% of patients with intermediate uveitis remained without specific diagnosis. 51.5% of all IU patients were diagnosed with pars plans; 4.9% of these cases were associated with MS, 3.6% with HCF with dense vitreous opacities, 1.5% with sarcoidosis and <1% with scleroderma or toxocarosis.

In the posterior uveitis group, a specific diagnosis could be established in 78.1% of patients. Ocular toxoplasmosis was present in 29%, followed by multifocal choroiditis in 17.7% and serpiginous choroidopathy in 4.8%. Posterior uveitis was associated with systemic infections in 5.8%, sarcoidosis in 2.5%, VKH in 1.8%, ocular toxocarosis in 1.4% and Behçet’s disease in 2.5% of cases.

For patients with panuveitis, a specific diagnosis could be established in 80.4% of cases. Panuveitis was associated with systemic infections in 26.7%, ocular toxoplasmosis in 15.8%, sarcoidosis in 7.1%, Behçet’s disease in 11.4%, VKH in 3% and acute retinal necrosis in 4.3% of cases.

Of the 2619 Uveitis cases, 39.4% stayed unclassified regarding specific diagnoses. The most predominant specific entity was HLA-B27–associated anterior uveitis (HLA-B27 AAU, with or without systemic disease), which was found in 19.5% of all patients, followed by ocular toxoplasmosis in 6.6%, herpetic uveitis in 7.1%, white dot syndromes (WDS) in 5.5%, systemic infections in 4%, HCF in 3.4%, sarcoidosis in 2.4%, Behçet’s disease in 1.9%, JIA in 2.2% and rheumatoid arthritis in 1.6% of the patients.

Table 2 summarizes the most common etiologies of uveitis cases and their anatomical distribution.

Age at onset of disease

The distribution of the patients’ age at disease onset is shown in Figure 2. The mean age at the onset of disease for the different specific entities can be seen in Table 3.

In the <7 years age group (n = 91, 3.45% of all patients), anterior uveitis represented the predominant anatomic group. The spectrum of specific diagnoses was narrow; JIA (29.6%), pars planets (10.3%), ocular toxoplasmosis (9.9%), sarcoidosis (2.2%), ocular herpes infection (6.6%), reactive arthritis (2.6%) and systemic infection (3.3%) were found, 35.1% of cases remained unclassified. In children below the ages of 7 and 17 years at the onset of disease (n = 215), an increase in the proportion of intermediate uveitis was noted (35.3%). Specific diagnoses were established in 53.9% of the cases. These included JIA (10.6%), ocular toxoplasmosis (14.4%), ocular herpes infection (3.7%), HLA-B27+ AAU

Table 1 Current systemic infections agents as causes of active uveitis

| Viral (n = 21) | Bacterial (n = 47) | Parasites (n = 7) | Fungal (n = 24) |
|---------------|-------------------|------------------|-----------------|
| VZV (n = 14)  | Mycobacterium tuberculosis (n = 21) | Ascites lumbricoides (n = 2) | Candida albicans (n = 22) |
| CMV (n = 5)   | Chlamydia trachomatis (n = 10) | Filaria (n = 2) | Aspergillus fumigatus (n = 2) |
| Other (n = 2) | Treponema pallidum (n = 9) | Giarda lamblia (n = 2) | Plasmodium falciparum (n = 1) |
| Borelia Burgdorferi (n = 7) | | | |

VZV: varicella virus infection; CMV: cytomegalovirus infection.
(3.7%), HCF (4.6%), Behçet’s disease (1.8%), ocular toxocarosis (0.9%), systemic infections (1.7%) and sarcoidosis (3.2%), with serpiginous choroiditis, multifocal choroiditis, SLE and chronic polychondritis each observed in less than 1% of patients.

In two-thirds of the patients, disease onset occurred between 17 and 60 years of age. In this age group, the disease entities were diverse and are listed in Table 2.

Onset of disease after the age of 61 years was found in 13.5% of the patients; the largest group had anterior uveitis (63.1%). The diversity of specific entities was again reduced; 44.7% of patients were without specific diagnoses, 17.5% had ocular herpes infections, 6.2% had systemic infections, 8.4% had HLA-B27+ AAU, 2.8% had rheumatoid arthritis, 2.5% had ocular toxoplasmosis, 2.5% had WDS, 1.4% had serpiginous choroiditis, 2.2% had ocular toxocarosis, 1.9%
Table 3 Specific entities found in at least 5 patients and age at onset of disease

| Entity               | Mean | Min | Max | Main (yrs) |
|----------------------|------|-----|-----|------------|
| JIA                  | 10.8 | 1   | 29  | <17        |
| APMPPE               | 20.2 | 17  | 27  | 17-40      |
| Reactive arthritis   | 34.7 | 7   | 66  | >17        |
| SLE                  | 30.7 | 13  | 53  | 36-60      |
| Toxoplasmosis        | 29.2 | 1   | 79  | 17-40      |
| Crohn’s disease      | 34.7 | 18  | 64  | 17-40      |
| Reiter’s disease     | 35.5 | 13  | 64  | 17-35      |
| Ulcerative colitis   | 43.4 | 24  | 49  | 17-60      |
| HCF                  | 32.4 | 12  | 70  | <40        |
| Behçet’s disease     | 31.1 | 15  | 55  | <40        |
| Ocular toxocarosis   | 47.2 | 11  | 71  | >17        |
| HLA-B27 only eye     | 37.8 | 9   | 82  | 17-60      |
| Unclassified cases   | 39.5 | 2   | 87  | 17-60      |
| Psoriatic arthritis  | 39.8 | 23  | 49  | 36-60      |
| Ancylostomiasis       | 39.5 | 17  | 75  | 17-60      |
| MFC                  | 38.6 | 16  | 77  | 17-60      |
| MS                   | 35.9 | 14  | 77  | 17-40      |
| Sarcoiosis           | 40.1 | 5   | 78  | 17-60      |
| Serpiginosis         | 45.3 | 15  | 76  | >36        |
| ARN                  | 43.9 | 7   | 62  | 1 >17      |
| Ocular herpes        | 47.8 | 2   | 88  | >36        |
| Systemic infections  | 47.5 | 3   | 80  | 17-60      |
| VKH                  | 42.6 | 30  | 78  | >30        |
| Birdshot             | 51.4 | 32  | 75  | 36-60      |
| MASQ                 | 55.2 | 10  | 80  | >36        |

For each classification, mean age of onset of disease (mean), minimum age at onset of disease (min), maximum age at onset of disease (max) and the main age group where this specific entity is found (main).

JIA: juvenile idiopathic arthritis; APMPPE: acute posterior multifocal placoid pigment epitheliopathy; SLE: systemic lupus erythematosus; HCF: Fuchs’ heterochromic cyclitis; MFC: multifocal choroiditis; MS: multiple sclerosis; ARN: acute retinal necrosis; VKH: Vogt-Koyanagi-Harada disease; MASQ: masquerade syndromes.

had sarcoidosis and <1% had either multifocal choroiditis, birdshot chorioretinopathy or reactive arthritis disease. There were no cases of Behçet’s disease and just one case of MS in the group with onset ages of over 60 years.

Discussion

In this study, we were able to establish specific diagnoses in 60.6% of 2619 patients; this is consistent with previous studies that reported ranges between 47 and 75% [29]. The relative frequencies of anatomical classifications are comparable to data published in other Middle European series from tertiary care centers but differ from the data of other Northern European countries, where up to 96% of the cases were reported to have anterior uveitis [30-35]. In the posterior and panuveitis groups, the rate of unclassified cases was the lowest at around 20%, and in the posterior uveitis group the frequency of toxoplasmosis was even higher than that among unclassified cases.

Overall, the largest diagnostic group comprised patients with ocular syndromes, followed by those with infectious diseases, arthritic diseases and, lastly, non-infectious systemic diseases.

When considering all patients, the major specific entities were HLA-B27 + AAU with or without systemic disease (19.5%), herpetic uveitis (7.1%) and ocular toxoplasmosis (6.6%).

The differential diagnosis of uveitis has changed over time. Tuberculosis and syphilis, the former main causes of uveitis, are now diagnosed in only 2.4% of patients [36,37]. More recently, however, increased frequency of these diseases has been noted. Factors that affect the changing patterns of uveitis include the rise of autoimmune diseases, appearance of new infections, description of new disease entities, better treatment of certain diseases, availability of new diagnostic tests and more refined classification of uveitis cases.

In the present study, the frequencies of different uveitis entities in different age groups were analyzed. In small children, only a few entities were discerned, including JIA, ocular toxoplasmosis, herpetic uveitis, pars planas, sarcoidosis, reactive arthritis, systemic infections and idiopathic anterior uveitis. With increasing age, the diversity of possible entities grew, reaching a peak between 30–40 years. In the elderly, the spectrum of specific diagnoses was again narrower; the major specific entities consisted of infections (herpes virus and toxoplasmosis) and masquerade syndromes. Some uveitis entities such as HCF, Behçet’s disease, and APMPPE appeared to affect adolescents or young adults with the greatest frequency; in children, JIA-associated Uveitis was most frequent. Nevertheless, most entities were observed in all age groups.

The high percentage of uveitis patients with systemic diseases and infections underpins the necessity of an interdisciplinary approach to uveitis therapy. Furthermore, rheumatologists and ophthalmologists have to be aware that a plethora of systemic autoimmune diseases and infections, as well as purely ocular syndromes, can cause uveitis. Infectious agents were involved in the cross a etiology of uveitis in almost 19% of all cases. This high percentage was mainly due to the high numbers of ocular toxoplasmosis (7.0%) and herpetic uveitis (7.1%) patients. These numbers imply that correct classification can have a decisive impact on the success or failure of therapies, and that immunosuppressive medications should not be given without ruling out infectious a etiology. However, the diagnostic quest should not lead to generalized and extensive investigations that generate unnecessary costs. The recommended course is a tailored approach in the hands of uveitis specialists and/or
rheumatologists/immunologists/infectiologists with uveitis experience. This would help to reinforce existing national guidelines, raise the standards of treatment, facilitate drug development, and shed light on the prognosis and course of the disease.

Abbreviations
ARN: Acute retinal necrosis; APM/PAPE: Acute posterior multifocal placoid pigment epitheliopathy; AU: Anterior Uveitis; CMV: Cytomegalovirus infection; HCF: Fuchs’ heterochromic cyclitis; JIA: Juvenile idiopathic arthritis; MS: Multiple sclerosis; MASQ: Masquerade syndromes; SLE: Systemic lupus erythematosus; SUN: Standardization of Uveitis nomenclature; VKH: Vogt-Koyanagi-Harada disease; VZV: Varicella virus infection; WDS: White dot syndrome.

Competing interests
The authors declare that they have no competing interests.

Authors’ contributions
The work presented here was carried out in collaboration between all authors. TBA and SMM defined the research theme. LM, WE, KM and HA analyzed the data and interpreted the results. TBA wrote the paper. All authors have contributed to, seen and approved the manuscript.

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References
1. Nussenblatt RB: The natural history of Uveitis. Int Ophthalmol Clin 1990, 10:303–308.
2. Gritz DC, Wong KE: Incidence and prevalence of uveitis in Northern California: the Northern California Epidemiology of Uveitis Study. Ophthalmology 2004, 111:491–500.
3. Bodaghi B, Cassoux N, Wechsler B, Hannouche D, Fardeau C, Papo T, Huong DL, Piette JC, LeHoang P: Chronic severe uveitis: etiology and visual outcome in 927 patients from a single center. Medicine (Baltimore) 2001, 80:263–270.
4. Gallagher MJ, Yilmaz T, Cervantes-Castañeda RA, Foster CS: The characteristic features of optical coherence tomography in posterior uveitis. Br J Ophthalmol 2007, 91:1680–1685.
5. Cassoux N, Giron A, Bodaghi B, Tran TH, Baudet S, Davy F, Chan CC, Lehoang P, Merle-Béral H: IL-10 measurement in aqueous humor for screening patients with suspicion of primary intraocular lymphoma. Invest Ophthalmol Vis Sci 2007, 48:3253–3259.
6. Menecio V, Bond SK, Towler HM, Kuo NW, Bahafo B, Wilson AG, Lightman S: Cytokine gene polymorphisms involved in chronicity and complications of anterior uveitis. Cytokine 2006, 35:200–206.
7. De Groot-Mijnes JD, Rothova A, Van Loon AM, Schuller M, Ten Dam-Van Loon NH, De Boer JH, Schuurman R, Weersink AJ: Polymerase chain reaction and Goldmann-Witmer coefficient analysis are complimentary for the diagnosis of infectious uveitis. Am J Ophthalmol 2006, 141:313–318.
8. Muthiah MN, Michaelides M, Child CS, Mitchell SM: Acute retinal necrosis: a national population based study to assess the incidence, methods of diagnosis, treatment strategies and outcomes in the United Kingdom. Br J Ophthalmol 2007, 91:1452–1455.
9. Green LR, Pavan-Langston D: Herpes simplex ocular inflammatory disease. Int Ophthalmol Clin 2006, 46:27–37.
10. Huerova V, Sanchez MC, Canto LM: Post-streptococcal syndrome uveitis. Acta Ophthalmol Scand 2007, 85:223–228.
11. Kharallah M, Yahia SB, Ladjimi A, Messaoudi R, Zaraoui S, Attia S, Jerzawi S, Jellis B: Pattern of uveitis in a referral centre in Tunisia, North Africa. Eye 2007, 21:33–39.
12. Soheilian M, Heidari K, Yazdani S, Shahsavai M, Ahmadiane H, Dehghan M: Patterns of uveitis in a tertiary eye care center in Iran. Ocul Immunol Inflamm 2004, 12:297–310.
13. Smith RL, Baasina GS, de Vries J: Classification of 750 consecutive uveitis patients in the Rotterdam Eye Hospital. Int Ophthalmol 1993, 17:71–76.
14. Tran VT, Auer C, Guex-Crosier Y, Pittet N, Herbot CP: Epidemiological characteristics of uveitis in Switzerland. Int Ophthalmol 1994-1995, 18:299–298.
15. Gayton JS, Woods AC: Etiology of uveitis: a clinical study of 562 cases. Arch Ophthalmol 1941, 26:963–1018.
16. Hendery DE, Genstler AJ, Smith RE, Rao NA: Changing patterns of uveitis. Am J Ophthalmol 1987, 103:131–136.
17. Chang JH, Wakefield D: Uveitis: a global perspective. Ocul Immunol Inflamm 2002, 10:263–279.
18. Ruthnam SR, Nanperumalsamy P: Global variation and pattern changes in epidemiology of uveitis. Indian J Ophthalmol 2007, 55:173–182.
19. Wakefield D, Chang JH: Epidemiology of uveitis. Int Ophthalmol Clin 2005, 45:1–13.
20. Rothova A, Suttorp-van Schulten MS, Frits Trefers W, Kljistra A: Causes and frequency of blindness in patients with intraocular inflammatory disease. Br J Ophthalmol 1996, 80:332–336.
21. Suttorp-van Schulten MS, Rothova A: The possible impact of uveitis in blindness: a literature survey. Br J Ophthalmol 1996, 80:844–848.
22. De Smet MD, Taylor SRJ, Bodaghi B, Miserocchi E, Murray PI, Pleyer U, Zierhut M, Barisani-Asenbauer T, LeHoang P, Lightman S: Understanding uveitis: The impact of research on visual outcomes. Prog Retin Eye Res 2011, 30:452–470.
23. Bloch-Michel E, Nussenblatt RB: International uveitis study group recommendations for the evaluation of intraocular inflammatory disease. Am J Ophthalmol 1987, 103:234–235.
24. Jabs DA, Nussenblatt RB, Rosenbaum JT: Standardization of uveitis nomenclature for reporting clinical data. Results of the First International Workshop. Am J Ophthalmol 2005, 140:509–516.
25. Rothova A, Buitenhuys HJ, Meenken C, Blinkman CJ, Linssen A, Alberts C, Luwendijk L, Kljistra A: Uveitis and systemic disease. Br J Ophthalmol 1992, 76:137–141.
26. Becker MD, Rosenbaum JT: Essential laboratory tests in uveitis. Dev Ophthalmol 1999, 31:92–108.
27. Chang JH, McCluskey PJ, Wakefield D: Acute anterior uveitis and HLA-B27. Surv Ophthalmol 2005, 50:364–388.
28. Levinson RD, Rajalingam R, Park MS, Reed EF, Gjertson DW, Kappel PJ, See RF, Rao NA, Holland GN: Human leukocyte antigen A29 subtypes associated with birdshot retinochoroidopathy. Am J Ophthalmol 2004, 138:631–634.
29. Zierhut M: Die Diagnostik der Uveitis. In Uveitis: Differentialdiagnosen. Edited by Zierhut M, Stuttgart, Germany: Kohlhammer Verlag, 1993:15–33.
30. McCannel CA, Holland GN, Helm CL, Cornell PJ, Winston JV, Rimmer TG: Causes of uveitis in the general practice of ophthalmology. UCLA Community-Based Uveitis Study Group. Am J Ophthalmol 1996, 121:35–46.
31. Kljistra A, Rothova A, Baasema GS, Zaal MJ, Fortuin ME, Schweitzer C, Glasius E, de Jong PT: Computer registration of uveitis patients. Doc Ophthalmol 1987, 67:139–143.
32. Montensen KK, Sjolie AK, Goldschmidt E: Uveitis. Eine epidemiologische Untersuchung. Ber Dtsch Ophthalmol Ges 1981, 78:97–101.
33. Jakob E, Reuland MS, Mackensen F, Harsch N, Fleckenstein M, Lorenz HM, Max R, Becker MD: Uveitis subtypes in a German interdisciplinary uveitis center — analysis of 1916 patients. J Rheumatol 2009, 36:127–136.
34. Pävönsalo-Hietanen T, Tuominen J, Vahtoranta-Lehtonen H, Saari KM: Incidence and prevalence of different uveitis entities in Finland. Acta Ophthalmol Scand 1997, 75:76–81.
35. Päivönsalo-Hietanen T, Vahtoranta-Lehtonen H, Tuominen J, Saari KM: Uveitis survey at the University Eye Clinic in Turku. Acta Ophthalmol 1994, 72:505–512.

36. Chao JR, Khurana RN, Fawzi AA, Reddy HS, Rao NA: Syphilis: reemergence of an old adversary. Ophthalmology 2006, 113:2074–2079.

37. Cimino L, Herbrecht OP, Ardili R, Salvarani C, Bicardi L: Tuberculous uveitis, a resurgent and underdiagnosed disease. Int Ophthalmol 2009, 29:67–74.

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