

CONTINUING MEDICAL EDUCATION

Red Eye: A Guide for Non-specialists

Andreas Frings, Gerd Geerling, Marc Schargus

SUMMARY

Background: Red eye can arise as a manifestation of many different systemic and ophthalmological diseases. The physician whom the patient first consults for this problem is often not an ophthalmologist. A correct assessment of the urgency of the situation is vitally important for the planning of further diagnostic evaluation and treatment.

Methods: This review is based on pertinent publications retrieved by a selective literature search in PubMed in August 2016 as well as on the authors' own clinical and scientific experience.

Results: Primary care physicians typically see 4–10 patients per week who complain of ocular symptoms. Most of them have red eye as the major clinical finding. A detailed history, baseline ophthalmological tests, and accompanying manifestations can narrow down the differential diagnosis. The duration and laterality of symptoms (uni- vs. bilateral) and the intensity of pain are the main criteria allowing the differentiation of non-critical changes that can be cared for by a general practitioner from diseases calling for elective referral to an ophthalmologist and eye emergencies requiring urgent ophthalmic surgery.

Conclusion: The differential diagnosis of red eye can be narrowed down rapidly with simple baseline tests and targeted questioning. Patients with ocular emergencies should be referred to an ophthalmologist at once, as should all patients whose diagnosis is in doubt.

► **Cite this as:**

Frings A, Geerling G, Schargus M: Red eye—a guide for non-specialists. *Dtsch Arztebl Int* 2017; 114: 302–12. DOI: 10.3238/arztebl.2017.0302

Every physician has been faced with the symptom “red eye” (1–3). According to one study, eye problems are the reason for 2 to 3% of visits to primary care physicians and emergency facilities (1). This corresponds to an average of 4 to 10 patients every week (2), the majority of whom have clinically apparent red eye (3). In general emergency rooms or in hospitals without a consulting ophthalmologist, non-specialists are confronted with the challenging decision of whether an urgent ophthalmological referral is necessary or not (1, 2). In the case of a foreign body event or other injury the patient should see an ophthalmologist immediately; the same applies if the cardinal symptom is pain, vision loss, rock-hard eyeball, or corneal involvement. An Australian study found that 64% of patients with red eye treated by primary care physicians were diagnosed incorrectly. In 10% of these cases there were serious clinical consequences (4). Another study from the same country concluded that literature for non-ophthalmologists is in short supply (5).

Learning goals

After reading this article the reader should be familiar with:

- The most frequently occurring differential diagnoses of red eye,
- the relevant questions to ask the patient and the basic investigations, and
- how to proceed in order to assess the urgency of referral to an ophthalmologist for further examination.

History

In the normal eye the conjunctiva shows a network of very fine blood vessels and the underlying sclera is white. Hyperemia, vascular ectasia, or subconjunctival hemorrhage therefore stand out. The degree of pain and the extent, uni-/bilaterality, location, duration, and course of the reddening must be recorded, together with any

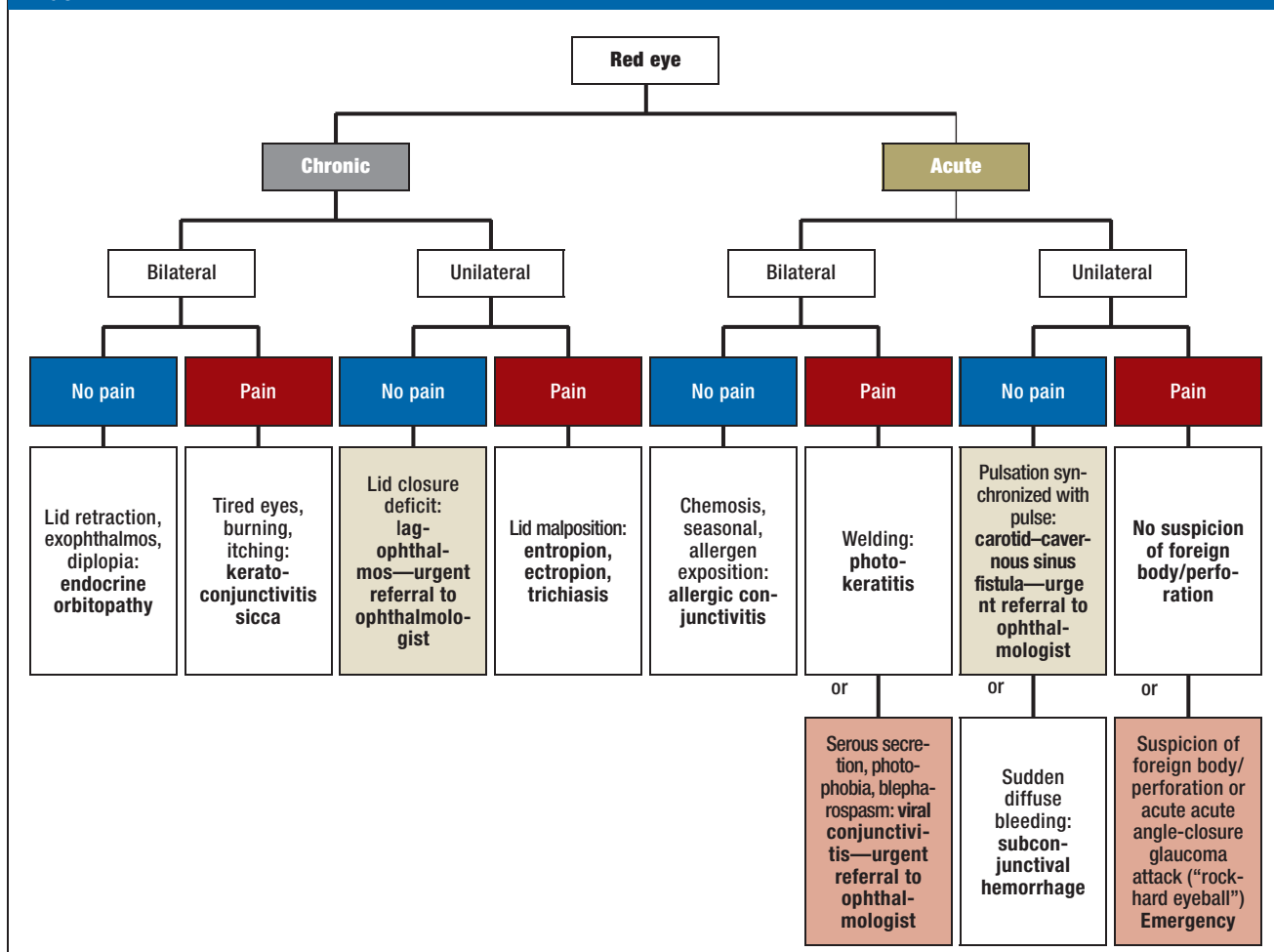
Department of Ophthalmology, Düsseldorf University Hospital: Dr. Frings, Prof. Geerling, FEBO, PD Dr. Schargus, MHBA, FEBO

Ophthalmological Medical Center Schweinfurt, Eye Hospital Schweinfurt-Gerolzhofen: PD Dr. Schargus, MHBA, FEBO

Incidence

According to one study, eye problems are the reason for 2 to 3% of visits to primary care physicians and emergency facilities.

FIGURE 1



Flow diagram for classification of red eye and its most frequently occurring causes

Flow diagram for differentiation and classification of causes of red eye for non-specialists. Bilateral findings often begin in one eye and affect the other eye later. “Emergency” means that immediate referral to an ophthalmologist is recommended. Figure modified from Cronau et al. (3)

accompanying symptoms. *Figures 1 and 2*, the *Table*, and *eTables 1 and 2* are intended to help the non-specialist arrive at the correct diagnosis.

Diagnostic work-up

The diagnostic work-up begins with ascertaining whether the red eye is an acute or chronic phenomenon. “Acute” is understood to mean duration of 7 days or less (*Figure 1*).

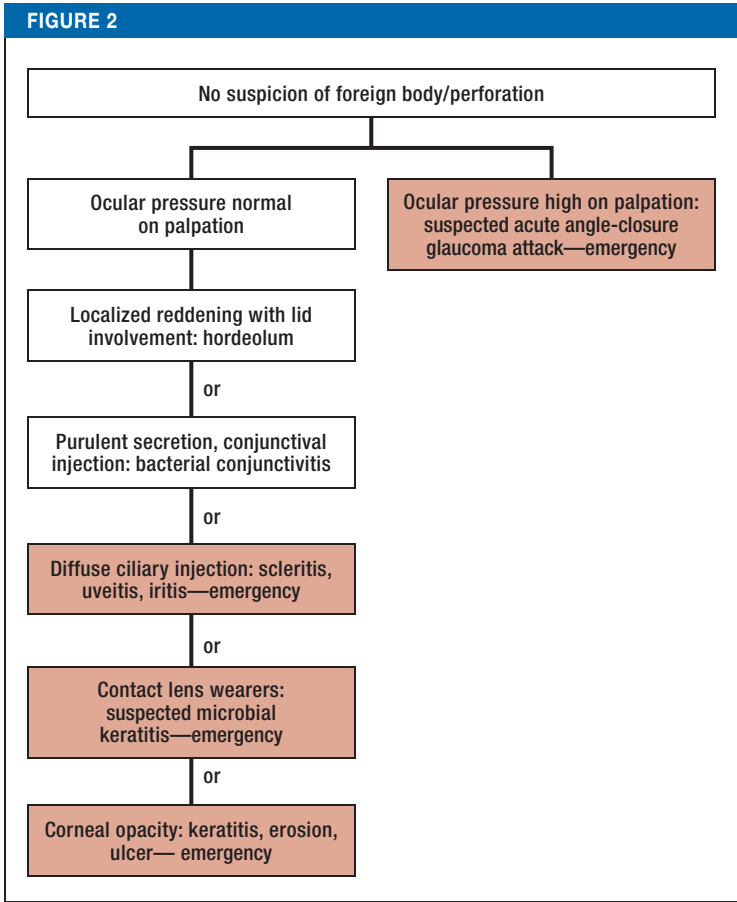
Both eyes should always be examined, as the laterality of red eye may be relevant to the diagnosis. The patient should also be asked about the occurrence of pain. Duration, laterality, and the presence and severity of pain are the cornerstones of diagnosis. In a patient with painful acute unilateral red eye, presence of a foreign body must be ruled out. Any patient with a positive history of a foreign body or trauma should be referred as an emergency to an eye hospital. If no foreign body is

Diagnosis

The duration and laterality of red eye and the presence and severity of pain are the cornerstones of diagnosis.

Diagnostic work-up

- Are both eyes affected?
- Is the red eye chronic or acute?



Principal causes of unilateral acute red eye, in the absence of suspicion of foreign body/perforation

suspected, the cornea must be inspected more closely. Superficial changes such as corneal erosion or deeper structural defects such as a corneal ulcer can often be discerned with the naked eye. If no such lesions are identified, the internal ocular pressure should be assessed by palpation, comparing the two eyes, to exclude an acute glaucoma attack (emergency). Should the internal ocular pressure be normal in a patient with painful acute unilateral red eye, one must consider inflammation of the sclera, uvea, or iris as a cause. Contact lens wearers who have painful acute unilateral red eye accompanied by diffuse ciliary injection and/or corneal opacity must be admitted for urgent treatment.

Conjunctivitis

The typical signs are red eye and eyelids stuck together on waking.

Painful acute bilateral red eye is often caused by viral conjunctivitis. As a rule the affected patients show a serous/clear secretion (overflow of tears = epiphora), photophobia, and blepharospasm. This also constitutes an ophthalmological emergency.

Any patient with non-painful acute unilateral red eye should be seen by an ophthalmologist within 24 h for exclusion of a carotid–cavernous sinus fistula. Urgent ophthalmological referral is also advisable for pain-free chronic unilateral red eye with lagophthalmos. Moreover, any patient with red eyes soon after ocular surgery should be referred to an ophthalmologist immediately.

Disorders of the cornea and conjunctiva

The term conjunctivitis refers to an inflammation of the conjunctiva alone, while keratoconjunctivitis is used to mean inflammation of the cornea and conjunctiva together. In both conjunctivitis and keratoconjunctivitis there is an inflammatory process of the ocular surface with widening of blood vessels (conjunctival injection), cellular infiltration, and watery, slimy, or purulent exudation (6). The typical signs are red eye and eyelids stuck together on waking; the eye may show glassy edema (chemosis) (Figure 3). The patient is particularly bothered by the sensation of a foreign body and a feeling of pressure, often in combination with photophobia and epiphora. If, in addition, an eyelid is affected by blepharospasm, corneal involvement is likely.

The many different possible causes of conjunctivitis (6) are divided into infectious and non-infectious factors. The latter include external irritants (smoke, dust, etc.), reduced production of tear fluid, eyelid malposition with mechanical irritation, and uncorrected refractive errors. The principal primary measure in the treatment of non-infectious conjunctivitis is avoidance of the triggering factor. Vasoconstrictive eye drops (active ingredient: tetrazone) should be prescribed for no more than a few days as a supportive measure, because they treat only the symptoms, not the cause, and may lead to tachyphylaxis, as described over 20 years ago in a case series of 11 patients (7). The opinion of an ophthalmologist should be obtained.

Infectious conjunctivitis is generally bacterial or viral. In the German-speaking area most cases of bacterial conjunctivitis are caused by staphylococci, streptococci, or pneumococci (6, 8, 9). Pain, pronounced reddening of the eye, and a mucopurulent secretion with formation of a yellowish crust are the typical signs. In principle, any

Vasoconstrictive eye drops

These should be prescribed for no more than a few days as a supportive measure, because they treat only the symptoms, not the cause, and may lead to tachyphylaxis.

available antibiotic can be considered for topical treatment of acute bacterial conjunctivitis, as randomized controlled trials and meta-analyses have demonstrated similar treatment effects (3, e1–e4). Two frequently used topical agents are gentamycin (an aminoglycoside antibiotic) and ofloxacin (a fluorochinolone antibiotic). The recommended minimum dosage is 3 × daily for 5 days. If no improvement is seen after 5 days, the patient should be referred to an ophthalmologist for urgent investigation.

Contact lens wearers

Contact lens wearers often suffer from a special form of bacterial (kerato-)conjunctivitis. For example, they may be infected with acanthamoebae or *Pseudomonas aeruginosa* (10, 11). Particularly with pseudomonads, hyperacute red eye with corneal ulceration may occur. The ulcer typically shows greenish-yellow mucus. The pathogens are frequently found in the contact lens fluid and/or container, and are often resistant to commonly used antibiotics. For timely diagnosis, the patient must provide the unwashed contact lens container together with the fluid and the lens itself and an ophthalmologist must obtain a direct swab from the focus of infection for microbiological analysis as part of an urgent ophthalmological consultation. Microbial keratitis can lead after just a few hours to corneal perforation and permanent visual loss or even to morphological loss of the eye (10). Keratoplasty “à chaud” (therapeutic corneal transplantation) with a prognosis of reduced visual acuity is then often the only possible form of treatment, as confirmed by a large retrospective case–control study published in 2015 (11). If corneal perforation is suspected, the eye should be protected by a non-compressive dressing with a compress secured at forehead and cheek.

Viral conjunctivitis

In contrast to bacterial conjunctivitis, bilateral red eye with a watery mucous secretion and itching should always arouse suspicion of a viral agent or an allergy. Viral conjunctivitis begins in one eye. Viral infection (most often by herpes simplex virus [12]) is accompanied by typical groups of herpetiform vesicles on reddened skin around the eye. In contrast to infection with varicella zoster virus, the skin lesions seen with herpes simplex do not respect the boundaries between dermatomes. Herpes simplex infection of the eye almost always represents a recurrence—the primary infection is often occult. Keratitis caused by fungi (keratomycosis)

TABLE

Common ophthalmological terms used to describe the findings (in alphabetical order)

Term	Meaning
Blepharospasm	Involuntary contraction/twitching of the eyelids
Chemosis	Glassy edema of the conjunctiva
Epiphora	Overflow of tear fluid
Halos	Colored rings around light sources
Injection	Vasodilatation with hyperemia
Lagophthalmos	Incomplete lid closure
Photophobia	Extreme sensitivity to light
Trichiasis	Chafing of the eyeball by inverted lashes

can only be confirmed by ophthalmological and microbiological investigation (e5).

Viral eye diseases such as epidemic keratoconjunctivitis are highly contagious (13, 14). They tend to be seasonal, caused by smear infection with adenoviruses, and manifest after an incubation period of 8 to 10 days (15). A patient with all the symptoms suffers severe pain in both eyes, accompanied by itching, light sensitivity, reddening, and tear overflow. Healing generally occurs after 2 weeks, usually with no consequences (15). Avoidance of contact is crucial: no handshakes, no sharing of the same flannel or towel, and strict observance of hygiene. All areas of the treating physician’s office entered by the patient must be decontaminated according to specific hygiene/disinfection standards. To prevent transmission, patients with red eye should be asked to wait in a separate room and should be given disposable gloves. The patient must strictly avoid contact with other people until the diagnosis is confirmed by an ophthalmologist and should not go to work. When a patient is referred for confirmation of the diagnosis, information should be provided in a brief telephone call. German law (§ 7, para. 1 of the Infection Protection Act) stipulates notification of the identity of the patient to the local health authority within 24 h of the direct demonstration of adenovirus in a conjunctival swab (e6). While epidemic keratoconjunctivitis is treated topically with

Contact lens wearers

Contact lens wearers often suffer from a special form of bacterial (kerato-)conjunctivitis. For confirmation of diagnosis, the unwashed contact lens container together with the fluid and the lens itself must be analyzed for pathogens.

Viral conjunctivitis

Viral eye diseases such as epidemic keratoconjunctivitis are highly contagious. They tend to be seasonal, caused by smear infection with adenoviruses, and manifest after an incubation period of 8 to 10 days.



Figure 3: Chemosis
Chemosis of the conjunctiva in infectious conjunctivitis



Figure 4: Subconjunctival hemorrhage
The arterial blood pressure should be measured to exclude a hypertensive crisis. If trauma cannot be ruled out, ophthalmosurgical exploration is required.



Figure 5: Corneal ulcer
White or gray discoloration of the cornea with an obvious tissue defect



Figure 6: Episcleritis
Inflammation of the connective tissue between sclera and conjunctiva with reddening, typically confined to one sector, and distinct dilatation of the episcleral vessels

moistening agents (15), herpetic (kerato-)conjunctivitis is treated topically and/or orally with a virostatic agent (e7–e10).

Non-infectious (kerato-)conjunctivitis

Non-infectious inflammation also requires timely treatment. Typical examples of non-infectious keratopathy are neurotrophic keratopathy and keratopathy due to lagophthalmos (see section entitled “Eyelid malposition”). In neurotrophic keratopathy the sensitivity of the cornea is greatly reduced, so that there is often a discrepancy between the patient’s subjective sensation of pain

and the actual findings (16). Patients typically have a history of surgery of the head (trigeminal neuralgia, acoustic neuroma), diseases that compromise the trigeminal nerve (tumor, brainstem hemorrhage), or infection with varicella zoster virus. Other possible causes are diabetes mellitus and multiple sclerosis. The treatment must include adequate moistening of the corneal surface with artificial tears and eye ointments (e.g., dexpanthenol) to avoid exposure keratopathy. Application of an occluding eye bubble moisture chamber at night, forming a moisture chamber, is still recommended. Potential new forms of treatment, e.g., topically applied nerve growth

Infection prophylaxis in the office

To prevent transmission, patients with red eye should be asked to wait in a separate room and to put on disposable gloves.

Topical treatment

While epidemic keratoconjunctivitis is treated symptomatically, herpetic (kerato-)conjunctivitis is treated topically and/or orally with a virostatic agent.

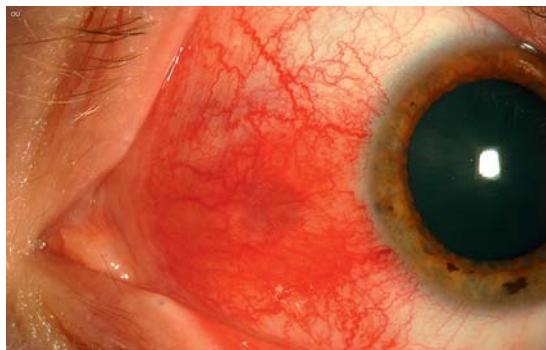


Figure 7: Scleritis

Typical appearance of scleritis: diffuse, “washed-out” red eye with dilatation of deep and superficial vessels



Figure 8: Hordeolum

Acute, painful bacterial inflammation of sebaceous or sweat glands at the margin of the eyelid



Figure 9: Lower-lid ectropion

External rotation of the lower lid, so that the tear fluid can no longer properly moisten the ocular surface. Conjunctival hyperemia owing to lagophthalmos is the result.



Figure 10: Carotid-cavernous sinus fistula

Bilateral massive dilatation of the conjunctival and episcleral vessels

factors, are currently being tested in randomized controlled trials (e11, e12).

Allergic reactions

Most case of acute allergic conjunctivitis are triggered by immunoglobulin-E-reactive antigens (type I: immediate-type allergy; classic “hay fever”). Chronic conjunctivitis, usually severe, is caused not by IgE but by T lymphocytes, which is why it is also sometimes referred to as a “cell-associated reaction” (type IV: delayed-type allergy). Examples of this type are vernal and atopic keratoconjunctivitis. The acute symptoms comprise bilateral swelling of the eyelids, epiphora,

itching, and conjunctival hyperemia with chemosis. Further exposure to the allergen should be avoided, and acute local and systemic antihistamines can be used. For long-term prophylaxis, mast cell stabilizers (sodium cromoglycate, lodoxamide) or H1 blockers of the second or third generation (e.g., olopatadine) can be prescribed, as shown in recently published reviews (17, e13). If the symptoms are severe, short-term local steroids (fluorometholone, loteprednol) can be given and a desensitization process initiated (18, 19). In the case of clinically meaningful type-IV sensitization after long-term use of eye drops containing preservative agents or sensitivity to ingredients of facial care

Non-infectious (kerato-)conjunctivitis

Typical examples of non-infectious keratopathy are neurotrophic keratopathy and keratopathy due to lagophthalmos.

Allergic reactions

The acute symptoms are bilateral swelling of the eyelids, epiphora, itching, and conjunctival hyperemia with chemosis.

products and cosmetics, avoidance of further contact with the allergen is mandatory. Systemic steroids or immunosuppressants can be used in the most severe cases. Epicutaneous testing to identify the allergen concerned should be arranged. As a rule the treatment of allergic reactions is interdisciplinary (18). Detailed information on dry eye disease can be found in the CME article by E.M. Messmer in issue 5/2015 of this journal (20).

Subconjunctival hemorrhage

Sudden subconjunctival hemorrhage (*Figure 4*) occurs particularly often in patients receiving oral anticoagulants and in those with poorly regulated or untreated arterial hypertension. The patients often report Valsalva maneuvers shortly beforehand (going to the toilet, gardening, etc.). In every “simple” subconjunctival hemorrhage, the patient’s blood pressure should be measured in the office to rule out an acute blood pressure crisis. The patient should be referred to an internist, because a large population-based study reported an elevated risk (hazard ratio 1.33) of (late) cardiovascular complications following subconjunctival hemorrhage (e14). The blood is resorbed spontaneously over a period of 2 to 4 weeks.

On the slightest suspicion of blunt or penetrating trauma, the patient must be transferred to an eye surgery center for operative exploration.

Corneal erosion/ulceration

Erosion of the cornea is a superficial defect of the epithelium. Pain, foreign body sensation, photophobia, and epiphora with blepharospasm are typical symptoms. The patients can usually describe the cause clearly.

If these symptoms occur after working with a drill or angle grinder, a metallic foreign body is often the cause. On the cornea, even after just a few hours in situ, such a fragment of metal leads to a rust ring that must be promptly removed by an ophthalmologist. Patients with no foreign body are referred to an ophthalmologist and can, if it seems advisable, be treated with an antibiotic eye ointment (e.g., chloramphenicol) in the meantime (e15). In the case of doubt, the patient should see an ophthalmologist without delay so that a penetrating injury can be ruled out. A special case is represented by red eye after surgery involving intubation anesthesia or in long-term ventilation on the intensive care unit. In these situations corneal erosion with reddening of the conjunctiva, usually in the lower third of the cornea, is caused by “dehydration” of the

cornea due to incomplete closure of the eyelids (21).

Deeper inflammation of the cornea can develop into keratitis or an ulcer (*Figure 5*). At this point there is a danger of corneal perforation. Therapeutic corneal transplantation often becomes necessary. Overall, the visual prognosis is unfavorable. Healing of the ulcer frequently leads to extensive scarring of the cornea and irregular astigmatism.

Photokeratitis

In photokeratitis the corneal inflammation is caused by ultraviolet rays. UV-C irradiation leads to acute necrosis of the epithelium over the exposed ocular surface. The typical situations include welding without goggles or a trip to a mountainous area without the necessary UV protection (e16, e17). Most patients present around 6 to 8 h after the event because the initially moderate, almost always bilateral pain and other symptoms (foreign body sensation, photophobia, epiphora) become unbearable and—due to the resulting blepharospasm—may lead to transitory functional blindness for up to 48 h. Although it is obvious what has happened, the patient should be seen by an ophthalmologist in a timely manner. The pronounced bilateral blepharospasm can be briefly interrupted by means of a single application of a topical local anesthetic (e.g., eye drops with oxybuprocaine hydrochloride 0.4%). There are no studies with high-level evidence, but oral analgesics should be given at the first contact with a physician, the patient should not rub his or her eyes, and vitamin A eye ointment should be used. The regeneration time of the damaged corneal epithelium is 24 to 48 h. Topical local anesthetics should not be prescribed.

Episcleritis

Episcleritis is a moderately painful inflammation of the connective tissue between the sclera and the conjunctiva with reddening usually confined to one sector and predominantly conjunctival injection (*Figure 6*). Scleritis, in contrast, is characterized by a markedly painful eyeball, often accompanied by impaired vision (22). Furthermore, the hyperemia in scleritis is not suppressed by the diagnostic (!) administration of a vasoconstrictor (*Figure 7*).

While episcleritis is generally triggered by exogenous factors (22), an underlying systemic disease (e.g., rheumatoid arthritis or vasculitis) can be demonstrated in up to 50% of patients with scleritis. In 20% of these cases, the systemic disease is of infectious

Subconjunctival hemorrhage

Sudden subconjunctival hemorrhage occurs particularly often in patients receiving oral anti-coagulants and in those with poorly regulated or untreated arterial hypertension. The patients often report Valsalva maneuvers beforehand.

Photokeratitis

In photokeratitis the corneal inflammation is caused by ultraviolet rays. UV-C irradiation leads to acute necrosis of the epithelium over the exposed ocular surface.

origin (e.g., varicella zoster virus). (22–24). The patient should always be urgently seen by an ophthalmologist. Episcleritis is usually self-limiting but may recur. Systemic comorbidity is rare. The treatment begins with moistening eye drops and non-steroidal anti-inflammatory drugs (NSARD), e.g., diclofenac (e18, e19). The treatment of scleritis ensues in stages (e18, e19). The underlying condition often has to be treated with anti-inflammatory drugs and immunosuppressants. Under certain conditions, new treatment strategies employing biologics/biosimilars such as tocilizumab or rituximab can be pursued (25, 26). Any delay in commencing treatment may result in a sharp increase in mortality from the underlying systemic disease (e20, e21). For instance, Foster et al. reported 10-year mortality of 30% in patients with rheumatoid arthritis (e22).

Intraocular disorders

Iritis and iridocyclitis (inflammation of the iris with involvement of the ciliary body and the anterior vitreous) are collectively referred to as anterior uveitis. Iritis is the most frequently occurring form of uveitis. In many cases no obvious causative factor is identified; nevertheless, immunological causes are more common than infections (e.g., syphilis or borreliosis) (27, 28). Rheumatic disorders, such as Behçet disease and Behçet syndrome, are particularly often associated with anterior uveitis (29). Furthermore, there is an association with inflammatory multisystem diseases such as sarcoidosis and juvenile idiopathic arthritis (30). The cardinal symptoms of iritis are photophobia, ciliary injection, and absence of the secretion typical of primary conjunctivitis.

Patients with a rheumatic disease or suspicion of anterior uveitis should be seen by an ophthalmologist within 24 h to avoid complications owing to exudative fibrinous inflammatory reactions. The treatment regimen for anterior uveitis depends on the etiology: frequent topical application of eye drops containing steroids, accompanied by pupil-dilating drops 2 or 3 times daily, is often sufficient (e23, e24).

Primary acute angle-closure glaucoma

Acute angle-closure glaucoma (“glaucoma attack”) is an acute ophthalmological emergency most often caused by pupillary block. The peripheral iris becomes attached to the cornea, leading to displacement of aqueous humor outflow with a resulting marked increase in internal ocular pressure (31).

Episcleritis

Episcleritis is a moderately painful inflammation of the connective tissue between the sclera and the conjunctiva with reddening usually confined to one sector and predominantly conjunctival injection.

Pain can radiate into the head and the teeth and also lead to secondary abdominal symptoms, cardiac rhythm disorders, and nausea via vagus stimulation. The cardinal symptom is the unilateral “rock-hard” eyeball, detectable by palpation of one eye and then the other: with the patient looking downwards, the eyeball is pressed down into the orbital fatty tissue with the index finger of one hand, holding it as still as possible, and the sclera is carefully indented with the index finger of the other hand. The deterioration in vision and the perception of colored rings around light sources (halos) are due to the corneal expansion (32). The increased internal ocular pressure causes intraocular ischemia and paralysis of the sphincter muscle of the pupil, so that the pupil is no longer round, moderately wide, and unresponsive to light. With the aid of a diagnostic pen light shone from the side, one can discern the flattening of the anterior chamber of the eye with approximation of the iris to the cornea.

Systemic and local treatment to lower the internal ocular pressure should be initiated without delay (e25). Many patients also require surgical intervention to deal with the angle closure (e26).

Disorders of the ocular adnexa

Hordeolum

Hordeolum (stye) occurs commonly in all age groups (1, 33). A usually acute bacterial inflammation of sebaceous or sweat glands at the margin of the eyelid manifests—in most cases painfully—as a raised red nodule (*Figure 8*). The exact incidence cannot be quantified (34). The recommended treatment comprises dry heat (e.g., from an infrared lamp) and an antibiotic ointment (e.g., gentamycin or ofloxacin, 3 × daily for a week), although no randomized clinical trials have been carried out. If there is no improvement, the patient should be referred to an ophthalmologist to exclude other possible diagnoses (e.g., sebaceous gland carcinoma) (1, 33).

Eyelid malposition

The common malpositions of the lower eyelids are ectropion and entropion, both usually a result of age-related slackening of the lid-retaining apparatus (35). In entropion the lid margin is inverted towards the eyeball and the eyelashes chafe against the surface of the eye (trichiasis). The danger is that defects of the corneal epithelium may arise, perhaps even a corneal ulcer. The test to determine whether entropion can be provoked,

Primary acute angle-closure glaucoma

Acute angle-closure glaucoma (“glaucoma attack”) is an acute ophthalmological emergency most often caused by pupillary block.

the patient should be asked to close the eye hard. In ectropion (*Figure 9*) the lower lid is rotated outwards and the tear fluid can no longer properly moisten the ocular surface, again resulting in defects of the cornea. One typical form of non-infectious keratopathy is lagophthalmic keratopathy, in which the eyelids close only incompletely or not at all. The most common cause is paralysis of the orbicular muscle of the eye in peripheral facial palsy.

In all of these situations the patient should be seen by an ophthalmologist. Surgical correction of the lid malposition is usually necessary.

Carotid–cavernous sinus fistula

A communication between the internal carotid artery and the cavernous sinus may occur spontaneously or result from trauma. In some cases the fistula is associated with bilateral massive, tortuous dilatation of the conjunctival and episcleral vessels (*Figure 10*). Further findings may include deterioration of vision, pulsatile exophthalmos, double vision, headache, and the perception of a rushing sound in the orbit that is audible on auscultation (36). The patient should be referred for urgent ophthalmological examination in an interdisciplinary center equipped to perform modern imaging modalities (e.g., magnetic resonance angiography).

Red eye in systemic diseases

Unilateral or bilateral red eye is frequently found in association with primarily systemic diseases. The systemic dermatological diseases in which red eye occurs include rosacea, molluscum contagiosum, and, commonly, severe atopy. Interdisciplinary management of these patients must not be neglected, as the ocular involvement may extend as far as sterile corneal perforation (37).

Conflict of interest statement

The authors declare that no conflict of interest exists.

Manuscript submitted on 18 August 2016, revised version accepted on 29 December 2016

Translated from the original German by David Roseveare

REFERENCES

1. Pflipsen M, Massaquoi M, Wolf S: Evaluation of the painful eye. *Am Fam Physician* 2016; 93: 991–8.

Lid malposition

In entropion the lid margin is inverted towards the eyeball and the eyelashes chafe against the surface of the eye. The danger is that defects of the corneal epithelium may arise, perhaps even a corneal ulcer.

2. Teo MA: Improving acute eye consultations in general practice: a practical approach. *BMJ Qual Improv Rep*. 2014; 3: pii: u206617.w2852.
3. Cronau H, Kankanala RR, Mauger T: Diagnosis and management of red eye in primary care. *Am Fam Physician* 2010; 81: 137–44.
4. Statham MO, Sharma A, Pane AR: Misdiagnosis of acute eye diseases by primary health care providers: incidence and implications. *Med J Aust* 2008; 189: 402–4.
5. O'Connor PM, Crock CT, Dhillon RS, Keeffe JE: Resources for the management of ocular emergencies in Australia. *Emerg Med Australas* 2011; 23: 331–6.
6. Azari AA, Barney NP: Conjunctivitis: a systematic review of diagnosis and treatment. *JAMA* 2013; 310: 1721–9.
7. Abelson MB, Butrus SI, Weston JH, Rosner B: Tolerance and absence of rebound vasodilation following topical ocular decongestant usage. *Ophthalmology* 1984; 91: 1364–7.
8. Alfonso SA, Fawley JD, Alexa Lu X: Conjunctivitis. *Prim Care* 2015; 42: 325–45.
9. Messmer EM: Keratitis—infectious or autoimmune? *Klin Monbl Augenheilkd* 2016; 233: 808–12.
10. Fernandes M, Vira D, Medikonda R, Kumar N: Extensively and pan-drug resistant *Pseudomonas aeruginosa* keratitis: clinical features, risk factors, and outcome. *Graefes Arch Clin Exp Ophthalmol* 2016; 254: 315–22.
11. Vazirani J, Wurity S, Ali MH: Multidrug-resistant *Pseudomonas aeruginosa* keratitis: risk factors, clinical characteristics, and outcomes. *Ophthalmology* 2015; 122: 2110–4.
12. Tsatsos M, MacGregor C, Athanasiadis I, Moschos MM, Hossain P, Anderson D: Herpes simplex virus keratitis: an update of the pathogenesis and current treatment with oral and topical antiviral agents. *Clin Experiment Ophthalmol* 2016; 44: 824–37.
13. Adhikary AK, Banik U: Human adenovirus type 8: the major agent of epidemic keratoconjunctivitis (EKC). *J Clin Virol* 2014; 61: 477–86.
14. Jhanji V, Chan TC, Li EY, Agarwal K, Vajpayee RB: Adenoviral keratoconjunctivitis. *Surv Ophthalmol* 2015; 60: 435–43.
15. Meyer-Rüsenberg B, Loderstädt U, Richard G, Kaulfers PM, Gesser C: Epidemic keratoconjunctivitis—the current situation and recommendations for prevention and treatment. *Dtsch Arztebl Int* 2011; 108: 475–80.
16. Chi JJ: Management of the eye in facial paralysis. *Facial Plast Surg Clin North Am* 2016; 24: 21–8.
17. Castillo M, Scott NW, Mustafa MZ, Mustafa MS, Azuara-Blanco A: Topical antihistamines and mast cell stabilisers for treating seasonal and perennial allergic conjunctivitis. *Cochrane Database Syst Rev* 2015; 6: CD009566.
18. Schröder K, Finis D, Meller S, Bühren BA, Wagenmann M, Geerling G: Seasonal and perennial allergic rhinoconjunctivitis. *Klin Monbl Augenheilkd* 2014; 231: 496–504.
19. Messmer EM: Ocular allergies. *Ophthalmologe* 2005; 102: 527–43.
20. Messmer EM: The pathophysiology, diagnosis, and treatment of dry eye disease. *Dtsch Arztebl Int* 2015; 112: 71–81.
21. Roth S, Thisted RA, Erickson JP, Black S, Schreider BD: Eye injuries after nonocular surgery. A study of 60,965 anesthetics from 1988 to 1992. *Anesthesiology* 1996; 85: 1020–7.
22. Sobolewska B, Zierhut M: Episkleritis und Skleritis. *Augenheilkunde up2date* 2015; 5: 51–62.
23. Diaz JD, Sobol EK, Gritz DC: Treatment and management of scleral disorders. *Surv Ophthalmol* 2016; 61: 702–17.
24. Dunn JP: Uveitis. *Prim Care* 2015; 42: 305–23.
25. Silpa-Archa S, Oray M, Preble JM, Foster CS: Outcome of tocilizumab treatment in refractory ocular inflammatory diseases. *Acta Ophthalmol* 2016; 94: e400–6.
26. Suhler EB, Lim LL, Beardsley RM, et al.: Rituximab therapy for refractory scleritis: results of a phase I/II dose-ranging, randomized, clinical trial. *Ophthalmology* 2014; 121: 1885–91.

27. D'Ambrosio EM, Cava M, Tortorella P, Gharbyia M, Campanella M, Iannetti L: Clinical features and complications of the HLA-B27-associated acute anterior uveitis: a meta-analysis. *Semin Ophthalmol* 2016; 12: 1–13.
28. Pathanapitoun K, Dodds EM, Cunningham ET Jr, Rothova A: Clinical spectrum of HLA-B27-associated ocular inflammation. *Ocul Immunol Inflamm* 2016; 18: 1–8.
29. Stavropoulos PG, Soura E, Kanelleas A, Katsambas A, Antoniou C: Reactive arthritis. *J Eur Acad Dermatol Venereol* 2015; 29: 415–24.
30. Heiligenhaus A, Minden K, Föll D, Pleyer U: Uveitis in juvenile idiopathic arthritis. *Dtsch Arztebl Int* 2015; 112: 92–100.
31. Wright C, Tawfik MA, Waisbourd M, Katz LJ: Primary angle-closure glaucoma: an update. *Acta Ophthalmol* 2016; 94: 217–25.
32. Schwenn O: Acute angle closure, chronic angle-closure glaucoma and their differential diagnoses. *Klin Monbl Augenheilkd* 2011; 228: 95–104.
33. Carlisle RT, Digiovanni J: Differential diagnosis of the swollen red eyelid. *Am Fam Physician* 2015; 92: 106–12.
34. Lindsley K, Nichols JJ, Dickerson K: Interventions for acute internal hordeolum. *Cochrane Database Syst Rev* 2013: CD007742.
35. Press UP: The aging eyelid. *Klin Monbl Augenheilkd* 2010; 227: 15–9.
36. Gonzalez Castro LN, Colorado RA, Botelho AA, Freitag SK, Rabinov JD, Silverman SB: Carotid-cavernous fistula: a rare but treatable cause of rapidly progressive vision loss. *Stroke* 2016; 47: 207–9.
37. Nivenius E, Montan P: Spontaneous corneal perforation associated with atopic keratoconjunctivitis: a case series and literature review. *Acta Ophthalmol* 2015; 93: 383–7.

Corresponding author

Dr. med. Andreas Frings
 Universitätsklinik für Augenheilkunde der
 Heinrich-Heine-Universität Düsseldorf
 Moorenstr. 5
 40225 Düsseldorf, Germany
 andi.frings@gmail.com

Supplementary material

For eReferences please refer to:
www.aerzteblatt-international.de/ref1717

eTables:

www.aerzteblatt-international.de/17m0302

Further information on CME

This article has been certified by the North Rhine Academy for Postgraduate and Continuing Medical Education. Deutsches Ärzteblatt provides certified continuing medical education (CME) in accordance with the requirements of the Medical Associations of the German federal states (Länder). CME points of the Medical Associations can be acquired only through the Internet, not by mail or fax, by the use of the German version of the CME questionnaire. See the following website: cme.aerzteblatt.de.

Participants in the CME program can manage their CME points with their 15-digit "uniform CME number" (einheitliche Fortbildungsnummer, EFN). The EFN must be entered in the appropriate field in the cme.aerzteblatt.de website under "meine Daten" ("my data"), or upon registration. The EFN appears on each participant's CME certificate.

This CME unit can be accessed until 25 June 2017, and earlier CME units until the dates indicated:

- „Aneurysmal Subarachnoid Hemorrhage“ (issue 13/2017) bis zum 25 June 2017,
- "ADHD" (Issue 9/2017) until 28 May 2017,

Please answer the following questions to participate in our certified Continuing Medical Education program. Only one answer is possible per question. Please select the most appropriate answer.

Question 1

Why should vasoconstrictive eye drops be prescribed as a supportive measure for no more than a few days?

- a) Because they lead to enduring blepharospasm
- b) Because they lead to photophobia
- c) Because they may lead to tachyphylaxis
- d) Because they lead to hyperemia
- e) Because they lead to increased internal ocular pressure

Question 2

What step should be taken immediately whenever bacterial conjunctivitis is suspected in a contact lens wearer?

- a) Treatment with local steroids
- b) A watch and wait strategy
- c) Prescription of new contact lenses
- d) Exclusion of accompanying viral disease
- e) Referral to an ophthalmologist

Question 3

With which disease is anterior uveitis most often associated?

- a) Multiple sclerosis
- b) Bekhterev disease
- c) Glaucoma
- d) Diabetes mellitus type 2
- e) Gout

Question 4

What are the typical symptoms of scleritis?

- a) Marked epiphora, sclera tending towards hyperemia, no accompanying systemic findings
- b) Slight, motion-dependent eyeball pain with purulent exudation
- c) Diffuse, "washed-out" red eye with severe eyeball pain
- d) Corneal opacity, usually pronounced circular corneal vascularization
- e) Tired, burning eyes, diplopia

Question 5

With what is spontaneous subconjunctival hemorrhage often associated?

- a) Diabetes mellitus type 2
- b) Intake of oral antihypertensives
- c) Existing arterial hypertension
- d) Perforating ocular injury
- e) Non-infectious conjunctivitis

Question 6

What should one do immediately in every case of simple subconjunctival hemorrhage?

- a) Examine the fundus of the eye
- b) Determine the visual acuity
- c) Check the visual field
- d) Administer an antibiotic ointment
- e) Measure the blood pressure

Question 7

What are the typical symptoms of an acute "glaucoma attack"?

- a) Rock-hard eyeball, purulent exudation, reduced visual contrast
- b) Soft eyeball, worsened vision, incipient myasthenia gravis
- c) Rock-hard eyeball, worsened vision, perception of colored rings around light sources
- d) Normal eyeball, worsened vision, restricted visual field
- e) Rock-hard eyeball, transient improvement in vision due to hyperemia, night blindness

Question 8

What should be the first step in red eye of unknown origin and suspicion of viral conjunctivitis?

- a) Ophthalmological assessment after the symptoms have worn off
- b) Microbiological exclusion of acanthamoebae
- c) First-line treatment with, as a rule, steroid eye drops
- d) Notification to the authorities within 48 h, even in isolated cases
- e) Put the patient in a separate waiting area, give him/her disposable gloves to wear

Question 9

A patient presents with painful acute unilateral red eye. The diagnostic work-up does not confirm the suspicion of a foreign body or perforation. The ocular pressure is high on palpation. What is one possible diagnosis?

- a) Uveitis
- b) Acute glaucoma attack
- c) Bacterial conjunctivitis
- d) Endocrine orbitopathy
- e) Seasonal chemosis

Question 10

A patient with known Behçet syndrome reports a 24-h history of severe eye pain on exposure to light. What is the first differential diagnosis that comes to mind?

- a) Posterior uveitis
- b) Cataract
- c) Glaucoma attack
- d) Anterior uveitis
- e) Photokeratitis

Supplementary material to:

Red Eye: A Guide for Non-specialists

Andreas Frings, Gerd Geerling, Marc Schargus

Dtsch Arztebl Int 2017; 114: 302–12. DOI: 10.3238/arztebl.2017.0302

eREFERENCES

e1. Behrens-Baumann W, Quentin CD, Gibson JR, Calthrop JG, Harvey SG, Booth K: Trimethoprim-polymyxin B sulphate ophthalmic ointment in the treatment of bacterial conjunctivitis: a double-blind study versus chloramphenicol ophthalmic ointment. *Curr Med Res Opin* 1988; 11: 227–31.

e2. Jauch A, Fsadni M, Gamba G: Meta-analysis of six clinical phase III studies comparing moxifloxacin 0.3% eye drops twice daily to five standard antibiotics in patients with acute bacterial conjunctivitis. *Graefes Arch Clin Exp Ophthalmol* 1999; 237: 705–13.

e3. Lohr JA, Austin RD, Grossman M, Hayden GF, Knowlton GM, Dudley SM: Comparison of three topical antimicrobials for acute bacterial conjunctivitis. *Pediatr Infect Dis J* 1988; 7: 626–9.

e4. Protzko E, Bowman L, Abelson M, Shapiro A: Phase 3 safety comparisons for 1.0% azithromycin in polymeric mucoadhesive eye drops versus 0.3% tobramycin eye drops for bacterial conjunctivitis. *Invest Ophthalmol Vis Sci* 2007; 48: 3425–9.

e5. Messmer EM: Keratitis—infectious or autoimmune? *Klin Monbl Augenheilkd* 2016; 233: 808–12.

e6. Robert Koch-Institut: Adenovirus-Konjunktivitis: RKI-Ratgeber für Ärzte. www.rki.de/DE/Content/Infekt/EpidBull/Merkblaetter/Ratgeber_Adenovirus_Konjunktivitis.html#doc2393452bodyText16 (last accessed on 3 January 2017).

e7. Tsatsos M, MacGregor C, Athanasiadis I, Moschos MM, Hossain P, Anderson D: Herpes simplex virus keratitis: an update of the pathogenesis and current treatment with oral and topical antiviral agents. *Clin Exp Ophthalmol* 2016; 44: 824–37. doi: 10.1111/ceo.12785

e8. Wilhelmus KR: Interventions for herpes simplex virus epithelial keratitis. *Cochrane Database Syst Rev* 2003; 3: CD002898. Update in: *Cochrane Database Syst Rev* 2007; 1: CD002898.

e9. Vrcek I, Choudhury E, Durairaj V: Herpes zoster ophthalmicus: a review for the internist. *Am J Med* 2017; 130: 21–6.

e10. Schuster AK, Harder BC, Schlichtenbrede FC, Jarczok MN, Tesarz J: Valacyclovir versus acyclovir for the treatment of herpes zoster ophthalmicus in immunocompetent patients. *Cochrane Database Syst Rev* 2016; 11: CD011503.

e11. Sacchetti M, Lambiase A: Diagnosis and management of neurotrophic keratitis. *Clin Ophthalmol* 2014; 8: 571–9.

e12. Yanai R, Nishida T, Chikama T, Morishige N, Yamada N, Sonoda KH: Potential new modes of treatment of neurotrophic keratopathy. *Cornea* 2015; 34: 121–7.

e13. Kam KW, Chen LJ, Wat N, Young AL: Topical olopatadine in the treatment of allergic conjunctivitis: a systematic review and meta-analysis. *Ocul Immunol Inflamm* 2016; 18: 1–15.

e14. Wang TJ, Keller JJ, Sheu JJ, Lin HC: A 3-year follow-up study on the risk of stroke among patients with conjunctival haemorrhage. *Acta Ophthalmol* 2013; 91: 226–30.

e15. Barequet IS, Harizman N, Ziv H, Rosner M: Healing rate of corneal erosions: comparison of the effect of chloramphenicol eye drops and ointment and high-concentration hyaluronic acid in an animal model. *Cornea* 2014; 33: 1080–2.

e16. Cullen AP: Photokeratitis and other phototoxic effects on the cornea and conjunctiva. *Int J Toxicol* 2002; 21: 455–64.

e17. Ting MA, Saha K, Robbie S: Mass photokeratitis following ultraviolet light exposure at a nightclub. *Cont Lens Anterior Eye* 2016; 39: 316–7.

e18. Tappeiner C, Walscheid K, Heiligenhaus A: Diagnosis and treatment of episcleritis and scleritis. *Ophthalmologe* 2016; 113: 797–810.

e19. Daniel Diaz J, Sobol EK, Gritz DC: Treatment and management of scleral disorders. *Surv Ophthalmol* 2016; 61: 702–17.

e20. Cocho L, Gonzalez-Gonzalez LA, Molina-Prat N, Doctor P, Sainz-de-la-Maza M, Foster CS: Scleritis in patients with granulomatosis with polyangiitis (Wegener). *Br J Ophthalmol* 2016; 100: 1062–5.

e21. Generali E, Cantarini L, Selmi C: Ocular involvement in systemic autoimmune diseases. *Clin Rev Allergy Immunol* 2015; 49: 263–70.

e22. Foster CS, Forstot SL, Wilson LA: Mortality rate in rheumatoid arthritis patients developing necrotizing scleritis or peripheral ulcerative keratitis. Effects of systemic immunosuppression. *Ophthalmology* 1984; 91: 1253–63.

e23. Jaffe GJ, Dick AD, Brézín AP, et al.: Adalimumab in patients with active noninfectious uveitis. *N Engl J Med* 2016; 375: 932–43.

e24. Thureau S, Pleyer U: Differential diagnosis of anterior uveitis. *Ophthalmologe* 2016; 113: 879–92.

e25. Wright C, Tawfik MA, Waisbourd M, Katz LJ: Primary angle-closure glaucoma: an update. *Acta Ophthalmol* 2016; 94: 217–25.

e26. Liebmann JM, Ritch R: Laser surgery for angle closure glaucoma. *Semin Ophthalmol* 2002; 17: 84–91.

eTABLE1

Classification of the causes of red eye by disease site

Characteristics	Diseases of the ocular surface									
	Conjunctivitis	Allergy	Dry eye	Subconjunctival hemorrhage	Erosion	Corneal ulcer	Photokeratitis	Episcleritis/scleritis		
	Bacterial	Viral								
Incidence	+++	+	+++	+	++	-	++	-		
(U)nilateral/ (B)ilateral	U/B	U/B	B	U	U	U	B	U		
Worsening of vision	+	+++	+	-	++	++	+++	+		
Photophobia	(+)	+++	(+)	-	+	+	+++	+		
(L)ocalized/ (D)iffuse reddening	D	D	D	L	D	D	D	L		
(S)uperficial/ (D)eep reddening	S	S	S	S	S	S	S	S/D		
Secretion: (P)urulent/ (W)atery/ (M)ucinous	P	W	MW	-	W	W/P	W	W		
Conjunctival hemorrhage	-	(+)	-	+	-	-	-	-		
Conjunctival chemosis	+	+++	-	-	-	-	+++	(+)		
Corneal lesions	(+)	-	+	-	+	++	+	-		
Foreign body event	-	-	-	+/-	+/-	-	-	-		
Elevated ocular pressure	-	-	-	-	-	-	-	-		
Further features	Gradual onset, often initially unilateral	Extreme symptoms, first on one side and then bilaterally	Chronic problem, often with a long history	Arterial hypertension, blood pressure crises, trauma, foreign body event	Trauma, foreign body, eyelid malposition, eyelid closure defect	Nerve palsy, eyelid malposition, eyelid closure defect	Welding in the previous 6-8 h	Frequently recurrent, deep injection of the sclera, investigate for systemic diseases		
Urgency of definitive diagnosis	Normal referral to ophthalmologist	Immediate referral to ophthalmologist following notification and hygiene measures	Normal referral to ophthalmologist	Normal referral to ophthalmologist	Urgent referral to ophthalmologist	Emergency, consult ophthalmologist immediately	Normal referral to ophthalmologist	Urgent referral to ophthalmologist		

Incidence: +++ very often, ++ often, + occasionally - rarely
Remainder of table: (+) may occur, +/- can occur with or without changes

eTABLE 2

Classification of the causes of red eye by disease site

Characteristics	Intraocular diseases		Diseases of the ocular adnexa	
	Iritis/ iritidocyclitis	Acute angle-closure glaucoma	Hordeolum	Carotid-cavernous sinus fistula
Incidence	–	–	++	–
(U)nilateral/ (B)ilateral	U	U	U	U
Worsening of vision	++	+++	(+)	+
Photophobia	++	++	–	–
(L)ocalized/ (D)iffuse reddening	D	D	+	D
(S)uperficial/ (D)eep reddening	S/D	S/D	Eyelid	D
Secretion: (P)urulent/ (W)atery/ (M)ucinous	–	–	P	–
Conjunctival hemorrhage	–	–	–	–
Conjunctival chemosis	(+)	–	–	(+)
Corneal lesions	–	+	–	–
Foreign body event	–	–	–	–
Elevated ocular pressure	–	+++	–	+
Further features	Rheumatic underlying disease	Halos, extreme pain, flattened anterior chamber, opaque cornea, unresponsive pupil; NB: general symptoms	Localized reddening and swelling of the eyelids	Conjunctival tortuosity, headache, rushing sound, pulsa- tile exophthalmos
Urgency of defini- tive diagnosis	Urgent referral to ophthalmologist	Emergency, immedia- te referral to eye sur- gery center	Normal referral to ophthalmologist	Urgent referral to ophthalmologist

Incidence: +++ very often, ++ often, + occasionally, – rarely

Remainder of table: (+) may occur, +/- can occur with or without changes