

A&I

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**Alternating hemiplegia
of childhood syndrome**

Williams syndrome

orphan**a**nesthesia

a project of the German Society
of Anaesthesiology and Intensive Care Medicine

SUPPLEMENT NR. 15 | 2021

OrphanAnesthesia –

ein krankheitsübergreifendes Projekt des Wissenschaftlichen Arbeitskreises Kinderanästhesie der Deutschen Gesellschaft für Anästhesiologie und Intensivmedizin e.V.

Ziel des Projektes ist die Veröffentlichung von Handlungsempfehlungen zur anästhesiologischen Betreuung von Patienten mit seltenen Erkrankungen. Damit will OrphanAnesthesia einen wichtigen Beitrag zur Erhöhung der Patientensicherheit leisten.

Patienten mit seltenen Erkrankungen benötigen für verschiedene diagnostische oder therapeutische Prozeduren eine anästhesiologische Betreuung, die mit einem erhöhten Risiko für anästhesieassoziierte Komplikationen einhergehen. Weil diese Erkrankungen selten auftreten, können Anästhesisten damit keine Erfahrungen gesammelt haben, so dass für die Planung der Narkose die Einholung weiterer Information unerlässlich ist. Durch vorhandene spezifische Informationen kann die Inzidenz von mit der Narkose assoziierten Komplikationen gesenkt werden. Zur Verfügung stehendes Wissen schafft Sicherheit im Prozess der Patientenversorgung.

Die Handlungsempfehlungen von OrphanAnesthesia sind standardisiert und durchlaufen nach ihrer Erstellung einen Peer-Review-Prozess, an dem ein Anästhesist sowie ein weiterer Krankheitsexperte (z.B. Pädiater oder Neurologe) beteiligt sind. Das Projekt ist international ausgerichtet, so dass die Handlungsempfehlungen grundsätzlich in englischer Sprache veröffentlicht werden.

Ab Heft 5/2014 werden im monatlichen Rhythmus je zwei Handlungsempfehlungen als Supplement der A&I unter www.ai-online.info veröffentlicht. Als Bestandteil der A&I sind die Handlungsempfehlungen damit auch zitierfähig. Sonderdrucke können gegen Entgelt bestellt werden.

OrphanAnesthesia –

a project of the Scientific Working Group of Paediatric Anaesthesia of the German Society of Anaesthesiology and Intensive Care Medicine

The target of OrphanAnesthesia is the publication of anaesthesia recommendations for patients suffering from rare diseases in order to improve patients' safety. When it comes to the management of patients with rare diseases, there are only sparse evidence-based facts and even far less knowledge in the anaesthetic outcome. OrphanAnesthesia would like to merge this knowledge based on scientific publications and proven experience of specialists making it available for physicians worldwide free of charge.

All OrphanAnesthesia recommendations are standardized and need to pass a peer review process. They are being reviewed by at least one anaesthesiologist and another disease expert (e.g. paediatrician or neurologist) involved in the treatment of this group of patients.

The project OrphanAnesthesia is internationally oriented. Thus all recommendations will be published in English.

Starting with issue 5/2014, we'll publish the OrphanAnesthesia recommendations as a monthly supplement of A&I (Anästhesiologie & Intensivmedizin). Thus they can be accessed and downloaded via www.ai-online.info. As being part of the journal, the recommendations will be quotable. Reprints can be ordered for payment.

Bisher in A&I publizierte Handlungsempfehlungen finden Sie unter:

www.ai-online.info/Orphsuppl
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Find a survey of the recommendations published until now on:

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orphananesthesia

Anaesthesia recommendations for Williams syndrome

Disease name: Williams syndrome

ICD 10: .I.

Synonyms: WS, Williams-Beuren syndrome

Disease summary: Williams syndrome (WS) results from the micro deletion of contiguous genes at chromosome 7q11.23 [1]. This genome part includes several genes [2] like ELN (ELastiN), encoding the tissue structural protein elastin [3], and Gtf2i (General Transcription Factor II-I), encoding a multifunctional phosphoprotein with roles in transcription and signal transduction [4]. This results in a multisystem disorder involving the cardiovascular [5], connective tissue and central nervous system. The incidence of typical forms is 1:20,000 people [6]. The clinical description associates a supravalvular aortic stenosis and/or a stenosis of the branches of pulmonary arteries, a psychomotor retardation and a facial dysmorphism [7,8]. An association with neonatal hypercalcaemia has also been reported [9]. The facial dysmorphism is characterised by an elfin facies syndrome: stellate pattern of the irises [10], broad forehead, periorbital fullness, high and rounded cheeks, pointed chin, flattened nasal bridge and upturned nose, long philtrum and wide and everted lower lip [11]. The most frequent orofacial alteration is malocclusion (94%) [12]. In addition to the psychomotor retardation, these patients have a disharmonious IQ (intelligence quotient) profile and are scoring better in verbal tests [13]; they are hypersocial and have major anxiety [14,15]. They also have musical abilities [16,17]. Cardiovascular abnormalities are common in WS [18]; most of them are arterial stenosis. Forty-five to 75% of WS patients suffer from supravalvular aortic stenosis and/or pulmonary arteries stenosis. Eighty percent of WS patients have cardiac abnormalities (aortic, mitral and/or tricuspid valve abnormalities, ventricular hypertrophy), they also have coronary artery abnormalities [19]. Systemic arteriopathy in other vessels of the body occur in 20% of cases, preferentially affecting the thoracic aorta (which can be associated with renal artery stenosis), but other sites are also possible (neck, limbs abdominal aorta...). Most of the cardiovascular abnormalities are found in the first year of life [11]. They also can suffer from endocrine dysfunction [20,21], pulmonary emphysema [2], ophthalmologic manifestation [22,23] or epilepsy [24].

Medicine is in progress



Perhaps new knowledge

Every patient is unique

Perhaps the diagnosis is wrong



Find more information on the disease, its centres of reference and patient organisations on Orphanet: www.orpha.net

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Typical surgery

Most of the surgical procedures in patients with WS occur during childhood [18]. Recent medical and surgical advancements have improved life expectancy and therefore both adult and paediatric patients with WS present for surgical procedures. Most common procedures include:

- Dental avulsion: hypoplasia of the enamel increase the prevalence of cavities [25],
- Scoliosis surgery [26,27],
- Strabismus surgery [22],
- MRI.

Type of anaesthesia

Both sedation and general anaesthesia can be performed; it is necessary to favour sedation whenever possible [28]. The risk of an adverse cardiac event during general anaesthesia is high: 11% [29]. Depending on age and surgery, peripheral locoregional anaesthesia seems to be a good choice. Spinal anaesthesia can be performed depending on favourable cardiologist evaluation.

Necessary additional pre-operative testing (beside standard care)

Evaluation by a cardiologist within one month of the procedure is recommended [28,30]. This evaluation should include [18]:

1. Physical examination with blood pressure taken in both arms and both legs. Echocardiography may not demonstrate a stenosis. Stenosis can be demonstrated with diminished pulses and measuring blood pressure in lower extremities [31].
2. ECG, in search of arrhythmias and/or QTc prolongation [32,33], history of ischaemia [19]. Consultation of an electrophysiologist, if needed.
3. Echocardiography: looking for cardiac abnormalities [19].
4. Screening for thyroid dysfunction [20,34] and diabetic disorders [21] and, in early childhood, looking for an hypercalcaemia.

Particular preparation for airway management

In the literature, WS facial abnormalities [12,28] are not associated with either difficult intubation or ventilation.

Particular preparation for transfusion or administration of blood products

No specific preparation is required for administration of blood products.

Particular preparation for anticoagulation

No specific preparation is required regarding anticoagulation.

Particular precautions for positioning, transportation and mobilisation

No specific precautions are required for positioning, transport or mobilisation.

Interactions of chronic disease and anaesthesia medications

Betablocker therapy should not be stopped.

Antiepileptic therapy should not be stopped.

Angiotensin Converting Enzyme Inhibitor should be stopped.

Anaesthetic procedure

Anaesthesia for patients with WS should be performed in a centre where a cardiologist and an intensivist and a (paediatric) cardiac intensive care unit are available. The risk of adverse cardiac events during general anaesthesia is high: 11% [29]. This risk may be reduced if the diagnosis is known before the surgery [35,36]. Patients at highest risk are those with supralvalvular aortic stenosis.

Information on preprocedural management have already been published [28] and include: hospital admission the night before the procedure, limitation of the period without fluid intake to two hours or begin of maintenance intravenous fluid, especially in patients under five years of age. Scheduling the case to be the first case of the day, thus limiting NPO time, is recommended.

Anxiety [15] is a specific characteristic of WS, it can be accompanied by tachycardia and must be considered as such. Pharmacological and non-pharmacological techniques must be used to reduce anxiety. Premedication is recommended to decrease anxiety. If not possible, patients with WS should be comforted by the presence of family or familiar guardian until loss of conscience. Particular attention must be provided for noise, as these patients are extremely sensitive to it [17]. The operating room must be quiet at the induction of anaesthesia [28].

Intravenous induction should be preferred to facial mask induction with sevoflurane [28,29]. Topical anaesthesia for venous catheter placement should be considered before anaesthesia. Experts [28] avoid propofol because of its haemodynamic effects which can be catastrophic in WS patients with supralvalvular aortic stenosis (hypotension, coronary ischaemia...). A combination of ketamine and midazolam seems to be a wise choice.

Some experts recommend to avoid succinylcholine because there is a theoretical risk of a hyperkalaemic response [34], but no such case has been published so far. Titrating neuromuscular blockade is accomplished by intraoperative monitoring of train of four.

It is important to mention the Kounis syndrome [37] that might occur in these patients and cause sudden death. The coronary lesions in Williams syndrome may be responsible for the occurrence of sudden death and/or cardiac arrest during anaesthesia [38–40].

Particular or additional monitoring

Monitoring of the electrocardiographic ST segment with a five lead electrocardiograph is recommended [28].

Four extremity blood pressure measurements should be done before and after anaesthesia/sedation; looking for an undiagnosed stenosis [28].

Haemodynamic goals: Maintenance of sinus rhythm at an age-appropriate heart rate, ensuring an adequate preload while avoiding rapid shifts in intravascular volumes, and aggressive treatment hypotension [41]. Invasive measurement of blood pressure can be considered depending on the patient's cardiac condition and type of surgical procedure. The target for blood pressure is the same as for patients without WS.

The anaesthesiologist should apply cardiovascular monitoring depending on the surgical procedure. A transoesophageal echocardiography is recommended for cardiac surgical procedures. Near-infrared spectroscopy can be helpful to estimate the tissue oxygenation of brain and somatic areas for both cardiac and non-cardiac procedures.

When monitoring the depth of anaesthesia with a specific tool (bi-spectral index, entropy, density spectral array...) it is strongly recommended to have an adequate level of anaesthesia and prevent vascular effects of the anaesthetic.

In early childhood, the risk of hypercalcaemia imposes regular evaluation of the serum calcium [9,28].

Possible complications

Patients with WS should be considered at risk for myocardial ischaemia [28].

WS patients have an increased risk of arrhythmias [32].

WS patients have an increased risk of cardiac arrest [29].

Post-operative care

WS patients should stay in the recovery room and the monitoring should be continued until recovery of near baseline cognitive and cardiovascular function. The presence of a family or familiar guardian should be encouraged. In case of a heavier and/or complicated procedure post-operative care must be pursued in the intensive care unit.

Disease-related acute problems and effect on anaesthesia and recovery

Disease triggered emergency-like situations are not common in WS.

Ambulatory anaesthesia

Ambulatory anaesthesia should not be considered.

Obstetrical anaesthesia

No data have been found in literature.

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