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Cough in the Athlete CHEST Guideline and Expert Panel Report



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BACKGROUND: Cough is a common symptom experienced by athletes, particularly after exercise. We performed a systematic review to assess the following in this population: (1) the main causes of acute and recurrent cough, either exercise-induced or not, (2) how cough is assessed, and (3) how cough is treated in this population. From the systematic review, suggestions for management were developed.

METHODS: This review was performed according to the CHEST methodological guidelines and Grading of Recommendations Assessment, Development and Evaluation framework until April 2015. To be included, studies had to meet the following criteria: participants had to be athletes and adults and adolescents aged ≥ 12 years and had to complain of cough, regardless of its duration or relationship to exercise. The Expert Cough Panel based their suggestions on the data extracted from the review and final grading by consensus according to a Delphi process.

RESULTS: Only 60 reports fulfilled the inclusion criteria, and the results of our analysis revealed only low-quality evidence on the causes of cough and how to assess and treat cough specifically in athletes. Although there was no formal evaluation of causes of cough in the athletic population, the most common causes reported were asthma, exercise-induced bronchoconstriction, respiratory tract infection (RTI), upper airway cough syndrome (UACS) (mostly from rhinitis), and environmental exposures. Cough was also reported to be related to exercise-induced vocal cord dysfunction among a variety of less common causes. Although gastroesophageal reflux disease (GERD) is frequent in athletes, we found no publication on cough and GERD in this population. Assessment of the causes of cough was performed mainly with bronchoprovocation tests and suspected disease-specific investigations. The evidence to guide treatment of cough in the athlete was weak or nonexistent, depending on the cause. As data on cough in athletes were hidden in a set of other data (respiratory symptoms), evidence tables were difficult to produce and were done only for cough treatment in athletes.

CONCLUSIONS: The causes of cough in the athlete appear to differ slightly from those in the general population. It is often associated with environmental exposures related to the sport training environment and occurs predominantly following intense exercise. Clinical history and specific investigations should allow identification of the cause of cough as well as targeting of the treatment. Until management studies have been performed in the athlete, current guidelines that exist for the general population should be applied for the evaluation and treatment of cough in the athlete, taking into account specific training context and anti-doping regulations. CHEST 2017; 151(2):441-454

KEY WORDS: athletes; cough; evidence-based medicine; exercise; guidelines; sports

Summary of Suggestions

1. In adult and adolescent athletes (\geq 12 years of age) complaining of acute or recurrent cough, we suggest to initially evaluate for the most commonly reported causes of cough in this group such as asthma, exerciseinduced bronchoconstriction (EIB), respiratory tract infections (RTIs), upper airway cough syndrome (UACS) due to rhinosinus conditions, and environmental exposures related to the sport training environments (Ungraded, Consensus Based).

2. In adult and adolescent athletes (\geq 12 years of age) complaining of acute or recurrent cough, we suggest that pulmonary function tests, particularly bronchoprovocation challenges, and assessment of allergy to common airborne allergens be performed in the investigation of cough to identify common etiologies such as asthma and EIB, and to evaluate the effects of environmental exposures such as allergens, respiratory irritants, and pollutants (Grade 2B).

3. In adult and adolescent athletes (\geq 12 years of age) complaining of acute or recurrent cough, we suggest to proceed with a systematic investigation based on suspected cause(s) from initial clinical assessment, with specific attention to the athlete's particular sport and training environment and context (exercise related or not) in which cough occurs, to determine its etiology (Ungraded, Consensus Based).

4. In adult and adolescent athletes (\geq 12 years of age) complaining of cough, we suggest a treatment trial directed at the suspected causes of cough similarly to the general population, but taking into account the sport performed and training environment. The anti-doping regulations and potential side effects of medications that could interfere with training performances should be reviewed and considered when appropriate (Ungraded, Consensus Based). Remarks: Anti-doping regulations are provided by the World Anti-Doping Agency (WADA) (https://www. wada-ama.org). The World Anti-Doping Code is a document that brings consistency to anti-doping rules, regulations, and policies worldwide. It is updated annually and the Prohibited List identifies the substances and methods prohibited to athletes in and out of competition. The Global Drug Reference Online (Global DRO) provides athletes and support personnel with information about the prohibited status of specific substances based on the current WADA Prohibited List. The Global DRO provides specific information on products sold in the United Kingdom, Canada, and the United States. The Global DRO is created through a partnership between UK Sport, the Canadian Centre for Ethics in Sport, and the US Anti-Doping Agency (http://www.globaldro.com).

5. In adult and adolescent athletes (\geq 12 years of age) complaining of acute and recurrent cough, we suggest that investigators perform randomized control trials to assess the effects of disease-specific and/or environment-specific (eg, cold-air-induced cough) treatments on cough because there are minimal data on how to optimally treat cough in these groups (Ungraded, Consensus Based).

Cough is a symptom frequently reported by athletes and even more so in those involved in endurance sports.¹⁻⁹ Conditions known to be responsible for cough in the general population may also be responsible for cough in athletes.¹⁰⁻¹² However, in this population, cough may also be associated with the inherent characteristics of specific sports and their training environments. For example, exercise-induced cough is particularly prevalent in swimmers and winter-sports athletes.^{13,14} In winter athletes, cough following exercise was more frequently reported during winter (71%) than during summer (44%).⁹ Regarding other sports, a prevalence of exercise-induced cough was reported in 14% of football

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ABBREVIATIONS: DRO = Drug Reference Online; EIB = exerciseinduced bronchoconstriction; EIVCD = exercise-induced vocal cord dysfunction; GERD = gastroesophageal reflux disease; RTI = respiratory tract infection; UACS = upper airway cough syndrome; WADA = World Anti-Doping Agency

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players.¹⁵ In addition, the large ventilatory volumes required by intense exercise may increase lung penetration of allergens or pollutants and cause airway dehydration.^{10,12,16-25} Airway rewarming may also possibly modulate airway responses.

As part of the global effort of the American College of Chest Physicians (CHEST) to develop guidelines for the management of cough, it was decided to fill a gap in the literature by developing a clinical practice guideline on how to assess and treat cough in athletes. To do so, we performed a systematic review of the available publications on this topic to search for evidence to make specific recommendations or suggestions about identification of the cause of cough as well as the assessment and treatment of cough in this population. It was suggested that evaluation and management of cough in athletes should take into account many specific characteristics and special considerations for athletes, such as daily training, training environment, doping regulations, and frequent traveling. As each cause of cough might have a specific treatment, accurate diagnosis is essential to avoid using unnecessary medication.

Methods

Search Strategy

Systematic review of the literature was performed. Studies published in PubMed, Scopus, and the Cochrane Library were identified using the following search terms: (cough OR exercise induced asthma OR rhinitis OR sinusitis OR laryngitis OR GERD OR gastro-esophageal reflux OR gastroesophageal reflux OR "chronic eosinophilic bronchitis" OR bronchoconstriction OR "environmental exposures" OR "vocal cord" OR "upper airway diseases" OR allergies) AND (athlete OR athletes OR skiers OR skiing OR swimmers OR swimming OR runners OR rowing OR marathon OR Olympic OR "competitive sports") were identified. The search was limited to articles in French and English; it began with the initiation of these databases and ended in April 2015. to review. References of included studies were searched to identify additional relevant publications. The study characteristics and inclusion/exclusion criteria were selected from the patient problem or population, intervention, comparison, and outcome (PICO) elements for all key clinical questions²⁶ (Table 1). To be included, studies had to meet the following criteria: subjects were described as athletes, adults and adolescents aged ≥ 12 years who complained of acute or recurrent cough, regardless of the duration and relationship to exercise. A subject was considered an athlete if he/she participated in an organized team or individual sport and took part in regular sport competitions against other athletes. Ideally, the sport had to require some form of systematic and usually intense training, regardless of the level of competition or number of training hours.²⁷

Study Selection

Being aware that there were only a few clinical trials on cough in the athlete, we included all types of study designs in our list of articles

Data Extraction and Quality Assessment

Two authors (L-P. B. and J. T.) independently analyzed the titles and content of the abstracts recovered to assess inclusion criteria. From the selected titles and abstracts, full reports about potentially relevant studies were obtained, and the authors independently assessed

TABLE 1	Cough in Athletes PICO Questions: In Adult and Adolescent Athletes 12 Years of Age or Older, (1) What
	Are the Main Causes of Recurrent or Acute Cough? (2) How Is Acute or Recurrent Cough Usually
	Evaluated? (3) How Should Recurrent and Acute Cough Be Treated?

Category	Inclusion Requirements	No	Yes
Study design	Is the article an evaluation study that uses one of the following designs? Clinical trial Systematic review Prospective observational Case report	Exclude	Continue
Populations	 Does the article report on patient groups that have all of the following characteristics? ≥ 12 years of age Athlete Cough of any duration Cough related or not to exercise 	Exclude	Continue
Intervention	Does the article report on the causes of cough, the results of diagnostic evaluation for cough, and the treatments of cough?	Exclude	Continue
Outcomes	Does the article report on: Diagnostic accuracy (eg, sensitivity, specificity, positive predictive value, negative predictive value, validity, reliability, responsiveness, feasibility) Therapeutic efficacy (eg, change in clinical practice, impact on patient or provider decision-making) Patient outcome efficacy (eg, acceptability, quality of life, chest pain, depression, or anxiety)	Exclude	Continue

eligibility of the studies. Disagreements were discussed and resolved by consensus. A third author, R. S. I., was available to help resolve disagreements if necessary.

For randomized controlled trials, the reviewers independently assessed the risk of bias criteria using criteria in the *Cochrane Reviews*. The criteria used were random sequence generation (selection bias), allocation concealment (selection bias), blinding of participants and personnel (performance bias) to the study protocol, blinding of outcome assessment (detection bias), incomplete outcome data (attrition bias), and selective reporting (reporting bias).²⁸ Quality assessment of observational studies was also performed.²⁹

Guideline Framework

Grading of recommendations/suggestions was made according to the Grading of Recommendations Assessment, Development and Evaluation framework as adopted by CHEST.³⁰ This framework separates the process of rating the quality of evidence from that of determining the strength of the recommendation. The quality of

Results

Of 1,283 references identified by our search string and screened, 60 articles were included in the analysis. The process of study selection is outlined in Figure 1. Five articles reported cough prevalence, 46 articles reported causes of cough, three papers discussed cough assessment, and six papers were on the treatment of respiratory symptoms in athletes. Although there were five randomized clinical trials on the treatment of respiratory symptoms in athletes, only one specifically assessed cough. Although the systematic review revealed many records of causes of cough in athletes, very few specifically addressed how to evaluate or treat cough in the athlete. As data on cough in athletes were often hidden in a set of other data (respiratory symptoms), evidence tables were difficult to produce and were done only for cough treatment in athletes. In general, the studies were randomized placebo-controlled, double-blind trials, and four of six were considered to be at low risk of bias.

Key Clinical Question 1: In adult and adolescent athletes 12 years of age or older, what are the main causes of recurrent or acute cough?

Causes of Cough

Although there has been no formal study on the comparative prevalence of causes of cough in athletes, cough in the athlete was mainly associated with asthma and EIB in 15 reports.^{1,7,8,11,33-43} Other associations were reported with airway dehydration and environmental exposures in six studies,^{21-24,44,45} RTIs in five studies,⁴⁶⁻⁵⁰ UACS due to rhinosinus conditions in two studies,^{13,51} and exercise-induced vocal cord dysfunction (EIVCD) in two studies.^{52,53} Although it is very common in athletes, we found no study on the

evidence was based on risk of bias, inconsistency, indirectness, reporting bias, and imprecision.³¹ The quality of evidence (ie, the confidence in estimates) was rated as high (A), moderate (B), and low or very low (C). The strength of recommendation was determined based on the quality of evidence and the balance of benefits and harms, patients' values and preferences, and availability of resources.³² Level 1 represented strong recommendations in which benefits or harms clearly outweighed the other. Level 2 represented a weak recommendation in which it was not clear that the benefits outweighed the harms (or vice versa), and new research could change the direction or strength of these recommendations.

Developing Recommendations/Suggestions

To be included in this guideline, a recommendation or suggestion had to be voted on by 75% of the eligible members of the entire Cough Expert Panel and achieve ratings of "strongly agree" or "agree" by 80% of the voting panelists. No cough panel member was excluded from voting.

effects of GERD on cough in this population. Eleven manuscripts reported less common causes of cough: exercise-induced pulmonary edema including nonimmersion sports in two publications,^{20,54} immersion sports in five studies,⁵⁵⁻⁵⁹ high-altitude pulmonary edema in one publication,⁶⁰ as well as pulmonary embolism in one report,⁶¹ salt water aspiration syndrome in one study,⁶² and passive smoking in one case.⁶³ No publication was found on heart conditions that induced cough in athletes.

Practice Recommendations/Suggestions

1. In adult and adolescent athletes (≥ 12 years of age) complaining of acute or recurrent cough, we suggest to initially evaluate for the most commonly reported causes of cough in this group such as asthma, exercise-induced bronchoconstriction (EIB), respiratory tract infections (RTIs), upper airway cough syndrome (UACS) due to rhinosinus conditions, and environmental exposures related to the sport training environments (Ungraded, Consensus Based).

Key Clinical Question 2: In adult and adolescent athletes $(\geq 12 \text{ years of age})$, how is acute or recurrent cough usually evaluated?

Assessment of Cough

Although there were no systematic evaluations of cough specifically proposed for athletes, the publications suggested that a search for causes of cough in athletes should take into account the history and temporal relationships between cough and exercise, its sudden onset or recurrence, and the training environment.

One study evaluated the seasonal variability of cough reflex sensitivity to capsaicin in winter athletes.⁹ The



Figure 1 – Preferred Reporting Items for Systematic Reviews flow diagram for the study selection. EIVCD = exercise-induced vocal cord dysfunction; RTIs = respiratory tract infections; UACS = upper airway cough syndrome.

prevalence of cough within 1 hour after exercise was higher in athletes compared with nonathletes, and cough was reported more frequently during winter in athletes. However, cough was not associated with cough reflex hypersensitivity to capsaicin, bronchoconstriction, or airway inflammation.⁹

Although the Allergy Questionnaire for Athletes⁶⁴ can be used as a predictor of allergy in athletes,⁶⁵ self-reported respiratory symptom questionnaires have not been shown to be reliable predictors for the diagnosis of exercise-induced asthma and airway hyperresponsiveness.^{5,6,66}

Practice Recommendations/Suggestions

2. In adult and adolescent athletes (\geq 12 years of age) complaining of acute or recurrent cough, we suggest that pulmonary function tests, particularly

bronchoprovocation challenges, and assessment of allergy to common airborne allergens be performed in the investigation of cough to identify common etiologies such as asthma and EIB, and to evaluate the effects of environmental exposures such as allergens, respiratory irritants, and pollutants (Grade 2B).

3. In adult and adolescent athletes (\geq 12 years of age) complaining of acute or recurrent cough, we suggest to proceed with a systematic investigation based on suspected cause(s) from initial clinical assessment, with specific attention to the athlete's particular sport and training environment and context (exercise related or not) in which cough occurs, to determine its etiology (Ungraded, Consensus Based).

Key Clinical Question 3: In adult and adolescent athletes (≥ 12 years of age), how should recurrent and acute cough be treated?

Treatment of Cough

There were only six studies looking at the effect of treatments on cough in the athlete (Table 2). Usually, treatment targeted the suspected or confirmed causes, but the effects of therapy were usually noted for the whole spectrum of respiratory symptoms and not only cough, or they were not specifically reported for cough. In these six studies, the effects of different treatments on respiratory symptoms were evaluated in 201 athletes, mostly young men (mean age, 20 years; 84% men). Medications assessed included ipratropium,⁶⁷ albuterol,⁶⁸ ciclesonide,⁶⁹ montelukast,⁷⁰ and budesonide.⁷¹ Compared with placebo, these drugs had modest or no effects on exercise-induced respiratory symptoms, including cough. Only a subgroup of crosscountry skiers experienced a decrease in cough frequency and perception of cough intensity following exercise with pretreatment with 80 mg of inhaled ipratropium.⁶⁷ Finally, in a randomized double-blind, placebo-controlled clinical trial, a daily supplement of 600 mg of vitamin C for 21 days was reported to lead to a reduction in the duration and incidence of symptoms of upper RTIs, including cough, before and following a 90-km run.⁷²

Practice Recommendations/Suggestions

4. In adult and adolescent athletes (\geq 12 years of age) complaining of cough, we suggest a treatment trial directed at the suspected causes of cough similarly to the general population, but taking into account the sport performed and training environment. The antidoping regulations and potential side effects of medications that could interfere with training performances should be reviewed and considered when appropriate (Ungraded, Consensus Based).

Remarks: Anti-doping regulations are provided by WADA. The World Anti-Doping Code is a document that brings consistency to anti-doping rules, regulations, and policies worldwide. It is updated annually and the Prohibited List identifies the substances and methods prohibited to athletes in and out of competition (see Table 3).

The Global DRO provides athletes and support personnel with information about the prohibited status of specific substances based on the current WADA Prohibited List. The Global DRO provides specific information on products sold in the United Kingdom, Canada, and the United States. The Global DRO is created through a partnership between UK Sport, the Canadian Centre for Ethics in Sport, and the US Anti-Doping Agency.

5. In adult and adolescent athletes (\geq 12 years of age) complaining of acute and recurrent cough, we suggest that investigators perform randomized control trials to assess the effects of disease-specific and/or environment-specific (eg, cold-air-induced cough) treatments on cough because there are minimal data on how to optimally treat cough in these groups (Ungraded, Consensus Based).

Discussion

There is a lack of evidence on the optimal management of cough in the athlete. Although we were able to find some observations about the main causes of cough in this population, there has been no systematic approach to its evaluation. Moreover, some causes of cough were not addressed in the publications screened. For example, despite its high prevalence in endurance athletes, ranging between 30% and 70%, no report looked at GERD and cough in this population.⁷³⁻⁷⁵ As GERD is indeed recognized to be a frequent cause of cough in the general population,⁷⁶ there is a need to look at this potential cause of cough in the athlete.

As asthma, EIB, and environmental exposure were the most common conditions associated with persistent cough, we suggest that pulmonary function tests, particularly bronchoprovocation challenges, and assessment of allergy to common airborne allergens should be considered in the investigation of cough in athletes. Investigation should, however, be guided by the probable cause of cough as determined by the initial clinical assessment (see Table 4).

Strengths and Limitations

Even though cough is a very commonly reported respiratory symptom in athletes, our analysis of the literature stresses the paucity of data on cough in the athlete. The strengths of this analysis include the extensive review of > 1,200 publications obtained according to the key words selected, the methodology used to perform the analysis, and its review by the CHEST Expert Cough Panel.

Its limitations are associated with the lack of highquality studies, the variable definition of an athlete among reports (resulting in a relatively heterogeneous group of athletes), the lack of specific analysis of cough among other types of respiratory symptoms, and the variable outcomes chosen to assess cough response to

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Author/Year Bordeleau et al ⁶⁷ / 2014	Study Design RCT	Symptoms Assessed Cough, breathless- ness, sputum, chest tightness, wheezing	Patients Cross-country skiers who reported exercise- induced respiratory symptoms	No. of Patients 22	Age (y) 17 ± 3	Sex (M/F) 10/12	Control Group (Nonathlete) No	Intervention (80 mg) vs placebo before cross- country skiing (3 min sprint preceded by 30-min warm-up)	Outcomes Number of cough (the number of coughs following exercise was recorded for 1 hour with microphone headsets), Borg score for symptom intensity and broncho- constriction (FEV ₁)	Efficacy No influence on the post- exercise maximal fall in FEV1, the frequency of cough following exercise and the perception of cough intensity. However, a subgroup of athletes seems to show beneficial response to ipratropium (decrease in number	WADA status Permitted	Quality of the evidence Moderate
Gong et al ⁶⁸ / 1988	RCT	Cough, sputum, substernal soreness, shortness of breath, wheezing, chest tightness, sore throat, nasal discharge	Endurance cyclists and triathletes without active asthma or the use of broncho- dilators or other medication	15	23 ± 5	14/1	No	Albuterol (180 mg) vs placebo, ozone exposure (0.21 ppm) or filtered air during ergocycle (40-km time trial, preceded by warm-up) in an environ- mental chamber	Ozone exercise- induced symptoms, broncho- constriction (FEV ₁ , FVC, FEF _{25-75%}), impairment of exercise performance (VE, Vo ₂ , RR, HR), and airways reactivity to inhaled histamine	The pre- treatment with albuterol did not significantly block or attenuate the effects of 0.21 ppm O ₃ on symptoms, exercise perfor- mance, lung function, and airways reactivity to histamine.	Permitted	Moderate

TABLE 2] Studies That Evaluated the Effect of Different Treatments on Respiratory Symptoms in Athletes

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TABLE 2] (Continued)

A 11 - 57		Symptoms		No. of		C (1)/D	Control Group	T.1		F.(C.	WADA	Quality of the
Author/Year Peters et al ⁷² / 1993	Study Design RCT	Assessed Common cold symptoms (runny nose, sneezing, sore throat, cough, and fever)	Patients Runners (90- km race) and nonathletic subjects who were not regular runners	Patients Athletes = 84 Control subjects = 73	Age (y) 5 < 25 y; $57 \ge$ 25 y and $<$ 40y; $22 \ge 40 y$ (athletes)	Sex (M/F) 82/2 (athletes)	(Nonathlete) Yes Similar age, not regular runners	Intervention 21 days before the race 600 mg of vitamin C daily (runners = 43; control group = 34) vs placebo containing citric acid (runners = 41, control group = 39)	Outcomes The incidence of post-race URT symptoms during post- race period	Efficacy The duration of the symptoms was shorter in the vitamin C supple- mented runners and their matched control subjects, decreased incidence of symptoms of URT before and for 3-wk post-race (running nose, sneezing, sore throat, cough, and fever), but the duration and the severity were less in control group receiving Vitamin C.	Permitted	Moderate
Koya et al ⁶⁹ / 2009	Retro- spective review	Respiratory symptoms at rest and on exercise	Regional athletes; junior and high-school students with asthma	39	18	28/11	No	Ciclesonide (200 or 400 µg/d) vs montelukast (10 mg/d) Duration of treatment: Ciclesonide (5.2 mo) Montelukast (5.6 mo)	The rate of improve- ment of the symptoms at rest and during exercise in accordance with the global assessment of efficacy,	Better symptom control and fewer changes of therapy in the Ciclesonide group, but not statistically significant.	Permitted	Low

(Continued)

TABLE 2] (Continued)

Author/Year	Study Design	Symptoms Assessed	Patients	No. of Patients	Age (y)	Sex (M/F)	Control Group (Nonathlete)	Intervention	Outcomes	Efficacy	WADA status	Quality of the evidence
									rate of change of therapy, pulmonary function test parameters, and FENO values	Significant improve- ment in % FVC, %FEV ₁ and %MMF in the Ciclesonide group in comparison with baseline values. Significant decrease of FENO values in the Ciclesonide group in comparison with baseline values.		
Helenius et al ⁷⁰ / 2004	RCT	Asthma-like symptoms (wheeze, cough, shortness of breath)	Ice hockey players from the Finnish Junior National League who reported at least 2 exercise- induced bronchial symptoms	16	18 ± 1	16/0	No	4 wk of montelukast 10 mg, 1 wk washout, and 4 wk placebo	The effect of montelukast on asthma- like symptoms, bronchial and responsive- ness and on indices of airway inflam- mation Daily diary on exercise- induced asthma-like symptoms (scale 0-10)	No difference between placebo and Montelukast in weekly or monthly scores for exercise- induced symptoms, especially cough. Montelukast had no effect on lung function or indices of airway inflam- mation in symp-	Permitted	Moderate

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(Continued)

Author/Year	Study Design	Symptoms Assessed	Patients	No. of Patients	Age (y)	Sex (M/F)	Control Group (Nonathlete)	Intervention	Outcomes	Efficacy	WADA status
										tomatic ice hockey players has compared with placebo.	
Sue-Chu et al ⁷¹ / 2000	RCT	Asthma-like symptoms (wheeze, abnormal breathless- ness, chest tightness)	Elite competitive cross- country skiers with 2 or more asthma-like symptoms on exertion, at rest, or on exposure to irritants	25	18 (16-20)	19/6	No	Inhaled budesonide 400 µg twice daily vs placebo Duration of treatment: budesonide (22.3 wk) placebo (22.8 wk)	The effect of budesonide during the competitive season on indices of inflam- mation in the bronchial mucosa and BAL fluid and on tenascin immuno- reactivity in the retirular	No difference between placebo and budesonide on airway inflam- mation, tenascin expression or clinical manifes- tations. Asthma-like symptoms were	Permitted

Quality of the evidence

Moderate

TABLE 2] (Continued)

 $FEF_{25-75\%} = \text{forced expiratory flow in the midexpiratory phase; } FENO = \text{fractional exhaled nitric oxide; } HR = \text{heart rate; } MMF = \text{maximum mid-expiratory flow; } O^3 = \text{ozone; } RCT = \text{randomized controlled trial; } RR = \text{respiratory rate; } URT = \text{upper respiratory tract; } VE = \text{ventilation (L/min); } Vo_2 = \text{oxygen consumption; } WADA = World Anti-Doping Agency.}$

basement

membrane

unchanged

in 17 (68%)

skiers.

Causes of Cough	Drug	WADA Status
Asthma	Inhaled β_2 -agonists	
	Salbutamol	Permitted ^a
	Formoterol	Permitted ^a
	Terbutaline	Prohibited
	Inhaled anticholinergic agents	Permitted
	Inhaled corticosteroids	Permitted
	Leukotriene antagonists	Permitted
	Oral corticosteroids	Prohibited ^b
RTIs	Antibiotics	Permitted
	First-generation antihistamines with or without oral decongestant	Permitted ^c
	Nasal ipratropium	Permitted
	Nasal corticosteroids	Permitted
	Codeine (short-term use)	Permitted
	Dextromethorphan	Permitted
UACS	Nasal corticosteroids	Permitted
	First-generation antihistamines with or without oral decongestant	Permitted ^c
	Second-generation antihistamines (nonsedative)	Permitted ^c
	Nasal ipratropium	Permitted
	Topical decongestants (eg, xylometazoline, phenylephrine)	Permitted
EIVCD	Proton pump inhibitors	Permitted

TABLE 3 Current Anti-Doping Status of Common Drugs Used for Cough in Different Conditions

It is essential always to check that the drugs that have been prescribed are permitted by the World Anti-Doping Agency on the site Global DRO at http:// www.globaldro.com. Prohibited drugs can be used by athletes only if a Therapeutic Use Exemption (TUE) is granted in accordance with the International Standards on TUE. The athlete should, however, list all medication used on the doping control form. RTIs = respiratory tract infections; WADA = World Anti-Doping Agency. See Table 3 legend for expansion of other abbreviations.

^aThe use of this drug is conditional; maximum daily doses of salbutamol and formoterol permitted are 1,600 and 54 μ g/d, respectively. A urinary concentration of salbutamol > 1,000 ng/mL or of formoterol > 40 ng/mL is considered by the WADA to be an adverse analytic finding unless the athlete proves the drug was used therapeutically. There will be a presumption that the substance was not taken by inhalation therapeutically, and the athlete will have the burden of demonstrating by means of a controlled pharmacokinetic study that the level found in the athlete's urine sample was the result of therapeutic inhaled use.

^bThese drugs are prohibited during competition only.

^cThe use of pseudoephedrine in competition is conditional; a maximum daily dose of 240 mg/d is permitted. The urinary concentration of pseudoephedrine should not be $> 150 \mu$ g/mL. Athletes must stop taking pseudoephedrine at least 24 hours before competition.

TABLE 4	Common	Causes of	f Cough	According to	o Sport	Training	Environment
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Sport Training Environment	Sport	Possible Stimuli	Causes of Cough
Outdoor winter sports	Cross-country skiing, biathlon, speed skating	Cold and dry air	Asthma, UACS, EIVCD
Indoor ice rink	Speed skating, figure skating, hockey	Nitrogen dioxide, other indoor pollutants	Asthma, UACS
Indoor pool	Swimming, water polo, diving, synchronized swimming	Chlorine derivatives	Asthma, UACS
Outdoor sports	Cycling, running	Ozone, sulfur dioxide, nitrogen dioxide, aeroallergens	Asthma, UACS, EIVCD
Indoor gymnasium	Track and field, combat sports	Aeroallergens	Asthma, UACS
Sea and lake	Open water swimming, triathlon, scuba diving	Immersion, cold, salt water	Pulmonary edema
Mountains, high altitude	Skiing, hiking	Cold, hypoxemia	Pulmonary edema

EIVCD = exercise-induced vocal cord dysfunction; UACS = upper airway cough syndrome.

treatment. In addition, the number of patients enrolled in the studies was small.

Future Research

Many gaps in knowledge need to be addressed regarding cough in the athlete. They include the following:

- Determination of the prevalence of specific causes of cough in athletes according to the type of sport
- Evaluation of the mechanisms underlying exerciseinduced cough in athletes
- Documenting the incidence of cough in exercise and bronchoprovocation challenges
- Determination of the optimal way to assess cough in the athlete
- Determination of the role of GERD as a cause of cough in the athlete
- Performance of randomized controlled trials to assess the effects of current treatment options for cough in the athlete in the presence or absence of an identifiable pathologic condition
- Performance of well-designed, high-quality studies to determine the benefits of therapies when specific causes of cough have been identified in the athlete, also taking into account the type of sport and environmental exposures

Conclusions

Cough appears to be a common symptom in the athlete, is often related to environmental exposures during training, and occurs predominantly following intense exercise. Clinical history and specific investigations should allow identification of the cause of cough and targeting of treatment. Because there is a lack of evidence about how to best investigate and manage cough in the athlete, current guidelines produced for the general population should be applied in the athlete, taking into account the training-specific context and anti-doping regulations.

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