The Cow's Milk-related Symptom Score **CoMiss**TM An awareness tool to recognize cow's milk-related symptoms in infants and young children



Background SYMPTOMS RELATED TO INTAKE OF COW'S MILK

Many infants present with symptoms that can be related to intake of cow's milk.

Cow's milk protein allergy (CMPA) however is often not considered as a diagnosis, mainly due to the lack of specific diagnostic marker.

There are two groups of patients:

- 1 those with obvious CMPA
- 2 those with unclear symptoms in different organ systems

The first group is easy to recognize:

- consists mainly of infants with immediate/rapid reactions
- most of the time IgE mediated

However, this group represents a minority of all the infants with suspected cow's milk-related symptoms seen by primary healthcare providers.

In many cases, parents report some or all of these symptoms:

- Infant crying, having 'colic'
- Difficult defecation or 2-3 loose stools per day
- Regurgitates 4 to 6 times a day
- Mild patches of atopic dermatitis

Due to the lack of diagnostic test for CMPA (other than the challenge test), an awareness tool to recognize cow's milk-related symptoms for use by primary healthcare providers may increase the awareness and the recognition of symptoms possibly related to cow's milk, leading to an accurate diagnosis at an earlier stage.



SYMPTOMS OF CMPA

The majority of affected children have one or more symptom involving one or more organ system, mainly the gastrointestinal tract or the skin.^{1,2}

The majority have at least two symptoms and symptoms in at least two organ systems.²

About 50%-70% have cutaneous symptoms, 50-60% gastrointestinal symptoms, and about 20-30% respiratory symptoms.³

- 1. Lifschitz C, Szajewska H. Cow's milk allergy: evidence-based diagnosis and management for the practitioner. Eur J Pediatr. 2015;174:141-50.
- 2. Høst A. Cow's milk protein allergy and intolerance in infancy. Some clinical, epidemiological and immunological aspects. Pediatr Allergy Immunol 1994; 5(5 Suppl): 1-36.
- 3. Vandenplas Y, Koletzko S, Isolauri E, Hill D, Oranje A, Brueton M, Staiano A Dupont C. Guidelines for the diagnosis and management of cow's milk protein allergy in infants. Arch Dis Child. 2007;92:902-8.

WHAT IS CoMiSS™?

- The CoMiSS™ awareness tool should be regarded as a tool to increase awareness of the most common symptoms of CMPA, that in turn can aid an earlier accurate diagnosis.
- The CoMiSS[™] should increase awareness and knowledge to minimize over- and under-diagnosis and to consider cow's milk-related symptoms.
- The CoMiSS[™] is also a tool that can be used to evaluate and quantify the evolution of symptoms during a therapeutic intervention.
- Published data using the CoMiSS[™] in clinical trials show that the predictive value of the tool is 80% if the score is >12 at the start and decreases to <6 within 2 weeks under an elimination diet with an extensively hydrolysed formula.

CoMiSS™ is not a diagnostic test for CMPA.

It does not replace a food challenge
and its usefulness will need to be evaluated
by a prospective randomized study.

WHO IS CoMISS™ INTENDED FOR?

The CoMiSSTM is intended for primary healthcare providers who are all too often very limited in time and therefore need a tool that is simple, fast and easy-to-use. It can be filled in by the healthcare provider or by the parents as preparation for the doctor's visit.

THE CoMiss™ AWARENESS TOOL

Most patients with obvious CMPA, such as those with anaphylactic reactions or immediate IgE mediated reactions will not be picked-up by the CoMiSS™.

However, many infants present with non-IgE mediated reactions to cow's milk protein. These infants are difficult to identify as they present with delayed reactions, often moderate to mild in severity and often in different organ systems.

Functional GI symptoms such as regurgitation and vomiting, constipation, crying and colic are considered as non-IgE mediated reactions. Atopic eczema is often not recognized as a possible consequence of ingestion of CMP.

Expert Consensus

ACTA PÆDIATRICA Acta Pædiatrica ISSN 0803-5253 A workshop report on the development of the Cow's Milk-related Symptom Score awareness tool for young children Yvan Vandenplas (yvan.vandenplas@uzbrussel.be)¹, Christophe Dupont², Philippe Eigenmann³, Ame Host⁴, Mikael Kuitunen⁵, Carmen Ribes-Koninck⁶, Neil Shah^{7,8}, Raanan Shamir⁹, Annamaria Staiano¹⁰, Hania Szajewska¹¹, Andrea Von Berg¹² Nell Shahir , Kadhan Shamir , Annamaria Sialano , rhama Szajewska , Angrea von Berg . 1.UZ Brussel, Free University of Brussels, Belgium . 2.Pediatric Gastroenterology, Hepatology and Nutrition Department, Necker Children's Hospital, Paris, France . 3.Pediatric Allergy Unit, University Hospitals of Geneva, Geneva, Switzerland . 4.Department of Paediatrics, Hars Christian Andersen Children's Hospital, Odense University Hospital, Odense, Denmark . 5.Phildren's Hospital, University of Helsinki and Helsinki University Central Hospital, Helsinki, Finland . 6.Paediatric Gastroenterology and Hepatology Unit. La Fe University Hospital, Valencia, Spain . 7.Great Ormond Street Hospital for Children London (UK) and KU Leuven, Belgium, UK . 3.TARCIID, Leuven, Belgium 7. Great Ormand Street Hospital for Children London (UK) and KU Leuven, Belgium, UK. 8. TARGID, Leuven, Belgium 9. Institute of Gastroenterology, Nutrition and Liver Diseases, Schneider Children's Medical Center of Israel, Sackier Faculty of Medicine Tel-Aviv University, Tel-Aviv, Israel 10. Department of Translational Medical Science, Section of Pediatrics, University of Naples Federico II, Naples, Italy 11. Department of Paediatric, The Medical University of Warsaw, Warsaw, Poland 12. Research Institute, Department of Pediatrics, Marien-Hospital, Wesel, Germany

atopic dematitis, awareness tool, constipation, cow's milk protein allergy, regurgitation

Correspondence Y Vandenplas, UZ Brussel, Department of Paediatrics, Laarbeeklaan 101, 1090 Brussels, Belgium. Tel: + 3224775780 |

Fax +3224775783 Email: war.vandenplas@uzbrussel.be

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Clinicians with expertise in managing children with gastrointestinal problems and/or atopic diseases attended a workshop in Brussels in September 2014 to review the literature and determine whether a clinical score derived from symptoms associated with the ingestion of cow's milk proteins could help primary healthcare providers. The Cow's Milk-related Symptom Score (CoMiSS), which considers general manifestations, dermatological, gastrointestinal and respiratory symptoms, was developed as an awareness tool for cow's milk-related symptoms. It can also be used to evaluate and quantify the evolution of symptoms during therapeutic interventions, but does not diagnose cow's milk protein allergy and does not replace a food challenge. Its usefulness needs to be evaluated by a prospective randomised study.

Conclusion: The CoMiSS provides primary healthcare clinicians with a simple, fast and easy-to-use awareness tool for cow's milk-related symptoms.

How to use THE COMISS™ AWARENESS TOOL IN CLINICAL PRACTICE

CoMiSS[™] quantifies the number and severity of symptoms.

Suspicion of 'cow's milk-related symptoms' based on the presence of a combination of the following symptoms:

GENERAL DISCOMFORT

Persistent distress or colic (≥ 3 hours per day wailing/irritable) at least 3 days/week over a period of > 3 weeks.

GASTROINTESTINAL SYMPTOMS

Frequent regurgitation, vomiting, diarrhea, constipation (with/without perianal rash), blood in stool.

RESPIRATORY SYMPTOMS

Runny nose (otitis media), chronic cough, wheezing (unrelated to infection).

DERMATOLOGICAL SYMPTOMS

Atopic dermatitis, angioedema, urticaria unrelated to acute infections, drug intake manifestations.

CoMiSS™: Cow's Milk-related Symptom Score

Last name:	First name:	Age:	Date:

PURPOSE

The CoMiSSTM is a simple, fast and easy-to-use awareness tool for cow's milk-related symptoms. It increases awareness of the most common symptoms of cow's milk protein allergy (CMPA) that in turn can aid an earlier diagnosis. CoMiSSTM can also be used to evaluate and quantify the evolution of symptoms during a therapeutic intervention.

INSTRUCTIONS

If there is a suspicion of cow's milk-related symptoms, rate the observed/reported symptoms by choosing the most appropriate score for each type of symptom. Once completed, add the scores together and put the total in the box at the bottom of the scoring form.

SYMPTOM	SCORE			
Crying*	0 1 2 3 4 5 6	≤ 1 hour/day 1 to 1.5 hours/day 1.5 to 2 hours/day 2 to 3 hours/day 3 to 4 hours/day 4 to 5 hours/day ≥ 5 hours/day	SCORE	
Regurgitation	0 1 2 3 4 5 6	0 to 2 episodes/day ≥ 3 to ≤ 5 of small volume > 5 episodes of > 1 coffee spoon > 5 episodes of ± half of the feeds in <half continuous="" feeds="" of="" regurgitations="" small="" the="" volumes=""> 30 min after each feed Regurgitation of half to complete volume of a feed in at least half of the feeds Regurgitation of the complete feed after each feeding</half>		
Stools (Bristol scale)	4 0 2 4 6	Type 1 and 2 (hard stools) Type 3 and 4 (normal stools) Type 5 (soft stool) Type 6 (liquid stool, if unrelated to infection) Type 7 (watery stools)		
Skin symptoms	0 to 6	Atopic eczema HEAD-NECK-TRUNK ARMS-HANDS-LEGS-FEET Absent 0 0 Mild 1 1 Moderate 2 2 Severe 3 3	SCORE	
	0 or 6	Urticaria NO YES 6		
Respiratory symptoms	0 1 2 3	No respiratory symptoms Slight symptoms Mild symptoms Severe symptoms	SCORE	

^{*} Crying only considered if the child has been crying for 1 week or more, assessed by the parents, without any other obvious cause.

TOTAL SCORE

READING THE RESULT

The scoring ranges from 0 to 33. Each symptom has a maximal score of 6, except respiratory symptoms where the maximal score is 3.

If final score ≥ 12, the symptoms are likely cow's milk related. This could potentially be CMPA.

If final score < 12, the symptoms are less likely related to cow's milk. Look for other causes.

CMPA diagnosis can only be confirmed by an elimination diet followed by an oral food challenge.

Vandenplas, Y., Dupont, C., Eigenmann, P., Host, A., Kuitunen, M., Ribes-Koninck, C., Shah, N., Shamir, R., Staiano, A., Szajewska, H. and Von Berg, A. (2015), A workshop report on the development of the Cow's Milk-related Symptom Score awareness tool for young children. Acta Paediatrica. doi: 10.1111/apa.12902



Download the CoMiSS™ scoring form at: NestleHealthScience.com/CoMiSS



CRYING

- The duration of crying is evaluated.
 The longer the crying, the higher the score.
- At least > 3 days/week for 1 week or more, assessed by the parents, without any other obvious cause.
- Crying relates to colic. Colic is defined as > 3 hours crying/day for at least 3 days during 1 week and since at least 1 week.
- Up to 3 hours of crying/day accounts for a score of 3.

Crying*

SCORE

- 0 ≤ 1 hour/day
- 1 1 to 1.5 hours/day
- 2 1.5 to 2 hours/day
- 3 2 to 3 hours/day
- 4 3 to 4 hours/day
- 5 4 to 5 hours/day
- 6 ≥ 5 hours/day
- ► Age > 2 weeks, < 4 months
- * Should only be considered if the child has been crying for 1 week or more, assessed by the parents and without any other obvious cause.

Regurgitation

SCORE

- 0 to 2 episodes/day
- 1 \geq 3 to \leq 5 of small volume
- 2 > 5 episodes of >1 coffee spoon
- 5 episodes of ± half of the feed in < half of the feedings
- 4 Continuous regurgitations of small volumes > 30 min after each feed
- 5 Regurgitation of half to complete volume of a feed in at least half of the feeds
- 6 Regurgitation of the complete feed after each feeding
 - ► Age > 2 weeks, < 6 months

REGURGITATION

- Using the score for grading the regurgitation.¹
- Rates the regurgitated volume and severity of the regurgitation.

STOOLS

Using the well established Bristol stool scale to evaluate the consistency of the stools:²

- Type 1 and 2 indicate constipation
- Type 3 and 4 are 'ideal consistencies'
- Type 5 soft stools
- Type 6 tends to indicate diarrhea
- Type 7 indicates overt diarrhea

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Type 1

Seperate hard lumps, like nuts (hard to pass)



• Type 2

Sausage-shaped but lumpy



Type 3

Like a sausage but with cracks on its surface



Type 4

Like a sausage or snake, smooth and soft



Type 5

Soft blobs with clear-cut edges (passed easily)



Type 6

Fluffy pieces with raged edges, a mushy stool



Type 7

Watery, no solid pieces. Entirely liquid

Stools (Bristol scale

SCORE

- 4 Type 1 and 2 (hard stools)
- Type 3 and 4 (normal stools)
- 2 Type 5 (soft stool)
- 4 Type 6 (liquid stool, if unrelated to infection)
- 6 Type 7 (watery stools)
- For at least one week

Skin symptoms

SCORE

0 to 6 Atopic eczema

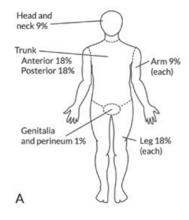
		HEAD-NECK TRUNK	ARMS-HAND LEGS-FEET
Abs	sent	0	0
Mil	d	1	1
Мо	derate	2	2
Sev	ere	3	3
or 6 Urt	icaria	NO	YES
		0	6

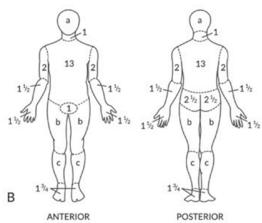
SKIN

 An easy to apply score based on an estimation of the surface covered by the dermatitis, using the surface estimation drawings from burns.

Estimated surface:

- ○ = 0
- < 1/3rd = 1
- 1/3-2/3rd **= 2**
- > 2/3rd = **3**





Relative percentage of body surface areas (% BSA) affected by growth

a —	½ of head
b-	1/2 of 1 thigh
-	1/2 of 1 lower lea

0 yr	1 yr	5 yr	10 yr	15 yr
91/2	81/2	61/2	51/2	41/2
23/4	31/4	4	43/4	41/2
21/2	21/2	23/4	3	31/4

RESPIRATORY

- Respiratory symptoms are considered in the CoMiSS™ although they have been given less importance (lower weighting) because most of the time chronic cough, runny nose and even wheezing are caused by viral infections.
- However, respiratory symptoms can be caused by cow's milk.

Respiratory symptoms

SCORE

- No respiratory symptoms
- 1 Slight symptoms
- 2 Mild symptoms
- 3 Severe symptoms
- ► Chronic cough
- Runny nose
- Wheezing

CoMiSSTM the scoring

The CoMiSS™ score ranges from 0 to 33.

Each symptom has a maximal score of six, with the exception of respiratory symptoms (maximal score of three).

An arbitrary cut-off value of ≥12 was selected as the criterion to pick up infants at risk of CMPA. A score of 12 requires the presence of at least two severe symptoms and a score higher than 12 requires the presence of at least three symptoms and the involvement of two organ systems.



USING THE SCORING FORM

If there is a suspicion of cow's milk-related symptoms, rate the observed/reported symptoms by choosing the most appropriate score for each type of symptom.

Calculate the final score:

- Total ≥ 12: This arbitrary cut-off value indicates that the symptoms are likely cow's milk related. This could potentially be CMPA. However, a CMPA diagnosis can only be confirmed by an elimination diet followed by an oral food challenge.
- Total < 12: symptoms less likely related to cow's milk. Look for other causes.



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