

How can a STRONG recommendation be based on VERY LOW quality evidence?

CHILDREN'S ONCOLOGY GROUP

The world's childhood cancer experts

Background

- <u>Grading of Recommendations</u>, <u>Assessment</u>, <u>Development and <u>Evaluation</u> (GRADE) is a widely used rating system</u>
- Several COG-endorsed supportive care guidelines have used the GRADE approach

GRADE classifies

Recommendations as **STRONG or WEAK** and

Quality of the Evidence* as VERY LOW, LOW, MODERATE or HIGH

*Reflects the level of confidence that the estimate of an intervention's effect is correct



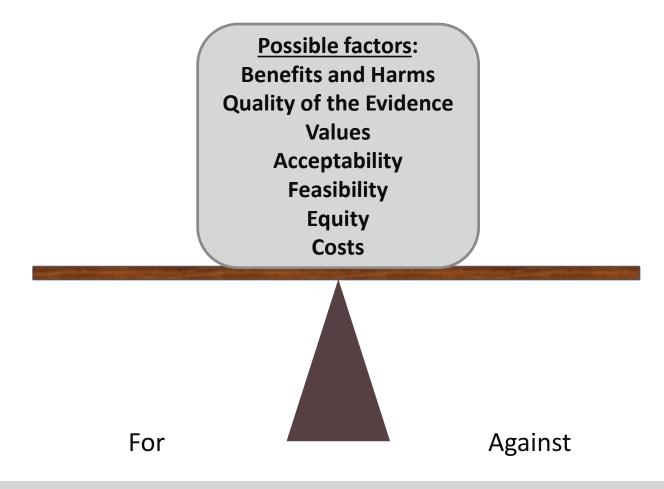
GRADE Quality of the body of evidence		
High ⊕⊕⊕	We are very confident that the true effect lies close to that of the estimate of the effect	
Moderate ⊕⊕⊕○	We are moderately confident in the effect estimate: The true effect is likely to be close to the estimate of the effect, but there is a possibility that it is substantially different	
Low ⊕⊕○○	Our confidence in the effect estimate is limited: The true effect may be substantially different from the estimate of the effect	
Very Low ⊕○○○	We have very little confidence in the effect estimate: The true effect is likely to be substantially different from the estimate of effect	
GRADE Strength of Recommendation		
Strong	Most people would want and should receive the recommended course of action. The recommendation can be adapted as a policy in most situations.	
Weak	There is a need for substantial debate and involvement of stakeholders in policy making. The majority of people would want the recommended course of action, but many would not. Clinicians should be more prepared to help patients to make a decision that is consistent with their own values.	

In this module we explain how a <u>STRONG</u> GRADE GUIDELINE RECOMMENDATION can be based on very low / low quality evidence and how to integrate such recommendations into your practice





GRADE recommendations: A balance of factors





Because GRADE incorporates separate judgments of evidence quality and strength of recommendation...

High quality evidence does not necessarily lead to a strong recommendation

and

Low quality evidence does not always lead to a weak recommendation

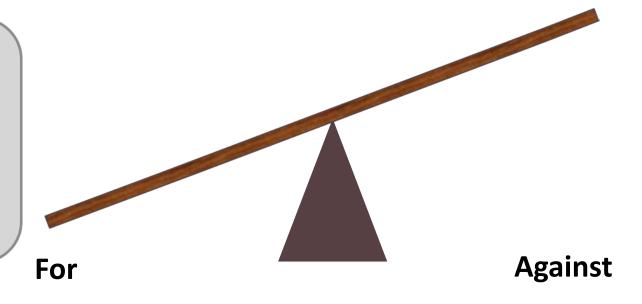




Strong recommendation

Possible factors:

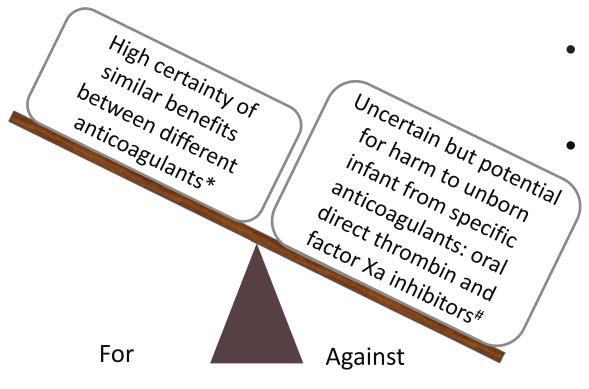
Benefits and Harms
Quality of the Evidence
Values
Acceptability
Feasibility
Equity
Costs



If after weighing all factors, the guideline panel is **very certain** that desirable effects outweigh undesirable effects, they will make a strong recommendation



GRADE example: Should oral direct thrombin and factor Xa inhibitors be used in pregnancy for women requiring anticoagulation?



Weighing the factors:

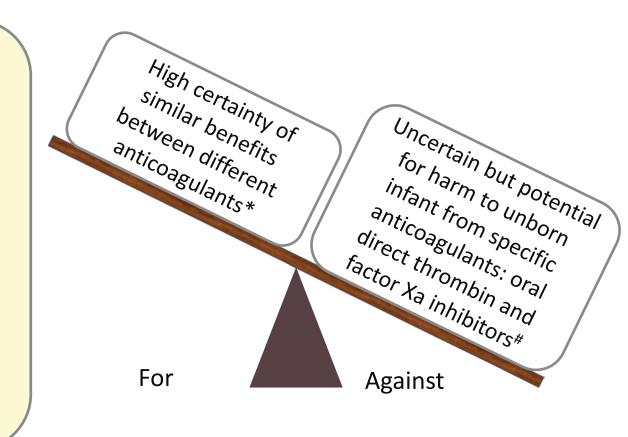
- *High certainty (i.e. high quality evidence)of similar effects of different anticoagulants
- # However, there is low quality evidence (i.e. low certainty of the effect) suggesting potential harm to the unborn infant with oral direct thrombin and factor Xa inhibitors



The guideline panel therefore made a **STRONG** recommendation against the use of oral direct thrombin and factor Xa inhibitors in pregnancy

This means:

Almost all women would not want to use oral direct thrombin and factor Xa inhibitors supplements in pregnancy and clinicians should not prescribe it in this setting







Implications for practice

Implications	GRADE Strong recommendation
For patients	Most individuals in this situation would want the recommended course of action, and only a small proportion would not.
	Formal decision aids are not likely to be needed to help individuals make decisions consistent with their values and preferences.
For clinicians	Most individuals should receive the intervention. Adherence to this recommendation according to the guideline could be used as a quality criterion or performance indicator.
For policy makers	The recommendation can be adopted as policy in most situations.

Source: http://www.gradeworkinggroup.org/





Want to learn more about GRADE?

Visit: https://cebgrade.mcmaster.ca/training/

Acknowledgement:

Example courtesy of Dr. Nancy Santesso, GRADE group

