

# Transforming the Healthcare Simulation Spectrum: **Now, Next and Beyond** 19 - 21 October 2022 Academia, Singapore



## **Effectiveness of non-pharmacological interventions (photobiomodulation**

versus oral cryotherapy) on oral-mucositis among patients undergoing chemotherapy conditioning prior to haematological stem cell transplantation

Chan Xin, Tay Lyn, Yap Shi Jed, Piyanee Yobas.

### **Background on oral-mucositis**

Oral-mucositis is a debilitating complication of chemo-conditioning regimens before haematopoietic stem cell transplantation (HSCT). Various health organisational guidelines recommended photobiomodulation (PBM) and oral cryotherapy (OC) as non-pharmacological oral-mucositis prevention for HSCT patients. However, current evidence is inadequate to assist clinicians in choosing the best-suited intervention for evidence-based individualized care plans.

## What is Photobiomodulation (PBM)?

Chemotherapy generates reactive oxygen species (ROS), which causes oxidative stress and tissue destruction, leading to oral-mucositis [1]. Thus, interventions that protect cells from chemo-induced ROS may aid in preventing oral-mucositis. At low concentrations, ROS are useful for normal cell signalling processes. At high concentrations, ROS are highly reactive molecules that can disrupt DNA bonds, and cause cell-mediated apoptosis. ROS are created by mitochondrial metabolism and are increased when the mitochondrial membrane potential (MMP) changes. MMP may be low due to oxidative stress, excitotoxicity, or electron transport inhibition. Photobiomodulation light absorption raises MMP to normal levels, lowering ROS generation [2]. It has been demonstrated that photobiomodulation photonic energy can reduce oxidative stress in oral-mucositis development [3].

## **Application and future implications**

Despite evidence showing similar effectiveness of PBM, current local practices only use OC in the management of oral-mucositis. There remained insufficient scientific evidence to determine which is superior. Therefore, future RCTs should conduct multicentre studies with large sample sizes that directly compare PBM versus OC effectiveness in preventing oral-mucositis among HSCT patients receiving chemo-conditioning. This is only possible with the application of PBM in real-life practices.

#### Implementation of PBM in Singapore

The PBM will be implemented in four hemato-oncological paediatric wards with evidence-based practice models to guide its execution. For PBM, the recommended parameters to reduce oral-mucositis severity and duration in HSCT patients before chemo-conditioning is 660-nm wavelength, 40mw power, 0.16J power, 1W/cm2 power density, 4J/cm2 energy density, and 0.04cm2 spot size [17]. For pain relief, the range of PBM total energy should be 15.36–20.16J [18].

Step 1

Step 3

Step 4

Synthesise

Evidence

Step 5

**Practice** 

Change



## **Comparing PBM vs OC**

Figure 1: Effectiveness of photobiomodulation and oral cryotherapy on severe oral-mucositis incidence.

	PBM/OC non-pharm int		Control		Risk Ratio		Risk Ratio		
Study or Subgroup	Events		Events	Total We	Weight	ht M-H, Random, 95% Cl		M-H, Random, 95% Cl	
1.2.1 PBM									
Antunes et al., 2007a	1	19	13	19	3.1%	0.08 [0.01, 0.53]	•		
Antunes et al., 2007b	3	19	17	19	6.0%	0.18 [0.06, 0.50]	•		
Djavid et al., 2011	0	27	3	28	1.7%	0.15 [0.01, 2.74]	•		
Ferreira et al., 2015	3	17	11	18	5.8%	0.29 [0.10, 0.86]	• •		
Hodgson et al., 2011a	12	20	9	20	8.0%	1.33 [0.73, 2.44]			
Hodgson et al., 2011b	12	20	11	20	8.3%	1.09 [0.64, 1.86]			
Hodgson et al., 2011c	11	20	7	20	7.5%	1.57 [0.77, 3.22]			
Hodgson et al., 2011d	4	20	2	20	4.0%	2.00 [0.41, 9.71]			
Khouri et al., 2009a	0	12	8	10	1.9%	0.05 [0.00, 0.77]	•		
Khouri et al., 2009b	0	12	5	10	1.8%	0.08 [0.00, 1.24]	•		
Salvador et al., 2017	0	27	4	24	1.7%	0.10 [0.01, 1.75]	•		
Silva et al., 2011	0	21	6	21	1.8%	0.08 [0.00, 1.28]	•		
Silva et al., 2015	0	20	4	19	1.7%	0.11 [0.01, 1.84]	•	<u>&gt;</u>	
Subtotal (95% CI)		254		248	53.2%	0.37 [0.17, 0.77]	-		
Total events	46		100						
Heterogeneity: Tau <sup>2</sup> = 1.	12; Chi <sup>2</sup> = 51.22, df	= 12 (P <	0.00001	); I <sup>2</sup> = 7	7%				

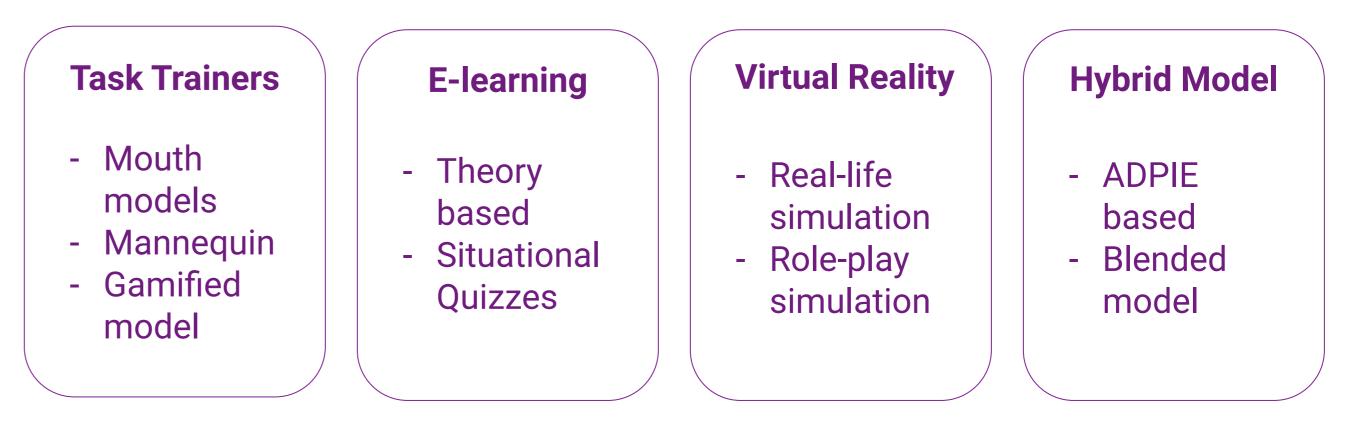
Identify	State the question/	Form a	Assemble,	Design &
issues		team	Appraise &	Pilot

#### **Utilising simulation in training**

purpose

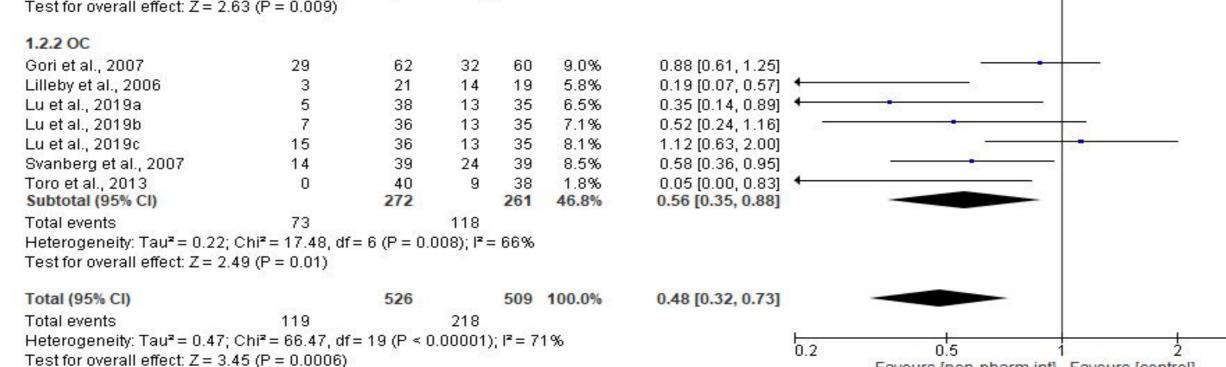
Step 2

Various methods will be used to train users.



#### Conclusion

With the data collected from the execution of PBM in Singapore, future research should explore the influence of chemo-conditioning intensities, patient-related risk factors and genetic-metabolic determinants of oral-mucositis on the severity of oral-mucositis.



Favours [non-pharm int] Favours [control]

#### **Results**

Test for subgroup differences: Chi<sup>2</sup> = 0.89, df = 1 (P = 0.35), l<sup>2</sup> = 0%

Meta-analyses included 18 RCTs (1018 HSCT patients). Both PBM and OC were effective in reducing oral-mucositis severity, severe oral-mucositis incidence, duration, and pain with small to large effect sizes. No significant differences were detected between PBM and OC across all outcomes. Subgroup analyses showed significant differences for chemo-conditioning regimens. The overall GRADE quality of evidence was low.

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