

A Modified Sidewall Adhesion Model: Hypothesis-Building and Testing Tool for a Surgical Experimental Research

Ospan A Mynbaev^{1,2,3,4}, Marina Yu Eliseeva², Andrea Tinelli⁵, Ioannis P Kosmas⁶, Antonio Malvasi⁷, Zhongjie Shi⁸, Mikhaylo V Medvediev⁹, Lyudmila V Lyutova¹⁰, Sergei V Pismensky^{11,12}, Zhomart R Kalzhanov¹³, Madina I Mazitova¹⁴, Tatyana I Babenko¹⁵, Michael Stark⁴

¹Department of Applied Mathematics, International Translational Medicine and Biomodeling Research Team, Moscow Institute of Physics & Technology (State University), Dolgoprudny, Moscow region, Russia

²The Department of Obstetrics, Gynecology & Reproductive Medicine, Peoples' Friendship University of Russia, Moscow, Russia

³Laboratory of Pilot Projects, Moscow State University of Medicine & Dentistry, Moscow, Russia

⁴The New European Surgical Academy, Berlin, Germany

⁵Department of Obstetrics and Gynaecology, Division of Experimental Endoscopic Surgery, Imaging, Minimally Invasive Therapy and Technology, Vito Fazzi Hospital, Piazza Muratore, Lecce, Italy

⁶Xatzikosta General Hospital, Ioannina, Ioannina, Greece

⁷Department of Obstetrics and Gynecology, Santa Maria Hospital, Bari, Italy

⁸Thomas Jefferson University, Philadelphia PA, USA

⁹Dnepropetrovsk State Medical University, Dnepropetrovsk, Ukraine

¹⁰Faculty of Biology, Moscow State University, Moscow, Russia

¹¹(Faculty of Basic Medicine, Moscow State University#), Moscow, Russia

¹²The V.I. Kulakov Research Center for Obstetrics, Gynecology and Perinatology, Moscow, Russia

¹³School of Health and Human Sciences, University of Essex, Colchester, UK

¹⁴The Department of Obstetrics & Gynecology N2, Kazan State Medical Academy, Kazan, Russian Federation

¹⁵The Department of Obstetrics & Gynecology, Stavropol State Medical Academy, Stavropol, Russia

Abstract

This editorial opinion article is based on our own experience and literature review. The NESA adhesion formation study team members recommend an unique model as a hypothesis building and testing tool in a surgical experimental research.

Body text

Postsurgical adhesions are believed to be the result of surgical tissue trauma in presence of multi factorial etiology of excessive healing, which may interfere with the physiological mechanisms at any stage [1]. The key points of postsurgical adhesions includes firstly the fixation of the intra-peritoneal structures to the parietal and/or visceral peritoneum, keeping them attached longer than in physiological wound healing [2] and secondly the initiation of excessive healing, caused by inflammation [3], which in turn may be triggered by surgical factors [4].

Postsurgical adhesions have evoked hard-to-cure and high-costly problems for the healthcare systems overall with such frequent complications such as small bowel obstructions with negative effects on patients' health, and increased workloads in clinical practice [5].

High-costs were associated with postsurgical adhesion complications as demonstrated in an analysis of annual inpatient care and expenditures based on a national hospital discharge data which were used to identify all abdominal adhesion related procedures performed in the United States in 1994 [6]: validated data with a ten-year follow-up of 12,584 patients undergoing lower abdominal surgery from the Scottish National Health Service Medical Record Linkage Database in UK [7] and in a study aimed to examine the natural course of patients following surgery for small bowel obstruction with a 14 year-follow-up in Sweden [8]. Under the evidence-based assumption that the risk of adhesions in colorectal surgery called for the necessity of opportunities and strategies for improvement the team of distinguished surgeons 'called for action' in their paper [9] to colorectal surgeons "to make prevention of adhesions more of a priority and importantly to inform patients of the risks associated with adhesion-related complications during the consent process".

From this view long lasting attempts of researchers and surgeons to find an optimal adhesion preventing adjuvant have legitimated their

expectations and the expenses were well worth it. Unfortunately until now most of suggested adhesion preventive adjuvant have shown their efficiency only in experimental models and failed in clinical trials.

Previously we demonstrated key steps of surgical research, namely the selection of the research questions, hypothesis building to set goals as well as the elaboration of a study design and interpretation of results based on our experimental studies and a systematic review of literature [10]. In our experimental researches we aimed to study an impact of surgical trauma and CO₂-insufflation on adhesion formation [3,10] to demonstrate the developing of research questions, the building of a hypothesis, and the setting of goals and the elaboration of a study design as well as the interpretation of results.

Therefore in this an editorial opinion paper we would like to demonstrate a rabbit sidewall adhesion model as a hypothesis building and testing tool. Because in the area of translational medicine and globalization an increased amount of information is required to apply state-of-the art methods based on results of evidence-based medicine in surgery.

***Corresponding author:** Ospan A Mynbaev, Department of Applied Mathematics, International Translational Medicine and Biomodeling Research Team, Moscow Institute of Physics & Technology (State University), Dolgoprudny, Moscow region, Russia, Tel: 79161271546; E-mail: spanmynbaev@crec.mipt.ru, ospanmynbaev@hotmail.com

Received October 10, 2013; **Accepted** November 06, 2013; **Published** November 11, 2013

Citation: Mynbaev OA, Eliseeva MY, Tinelli A, Kosmas IP, Malvasi A (2014) A Modified Sidewall Adhesion Model: Hypothesis-Building and Testing Tool for a Surgical Experimental Research. Clinics Mother Child Health 11: e101. doi: 10.4172/2090-7214.1000e101

Copyright: © 2014 Mynbaev OA, et al. This is an open-access article distributed under the terms of the Creative Commons Attribution License, which permits unrestricted use, distribution, and reproduction in any medium, provided the original author and source are credited.

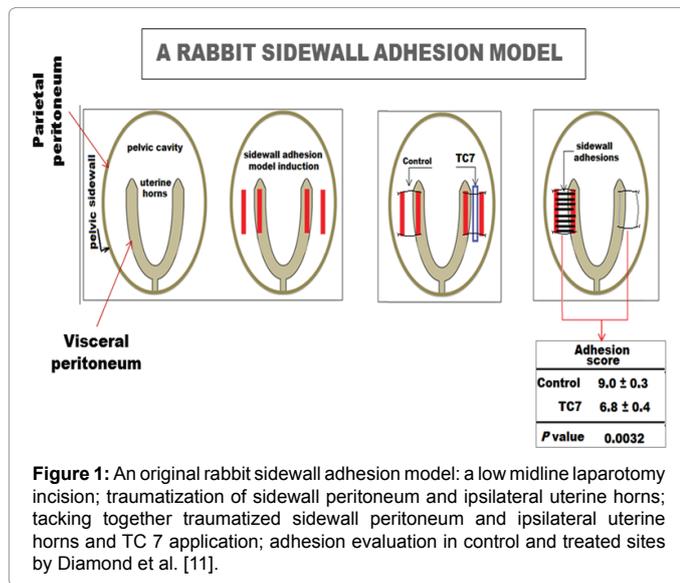


Figure 1: An original rabbit sidewall adhesion model: a low midline laparotomy incision; traumatization of sidewall peritoneum and ipsilateral uterine horns; tacking together traumatized sidewall peritoneum and ipsilateral uterine horns and TC 7 application; adhesion evaluation in control and treated sites by Diamond et al. [11].

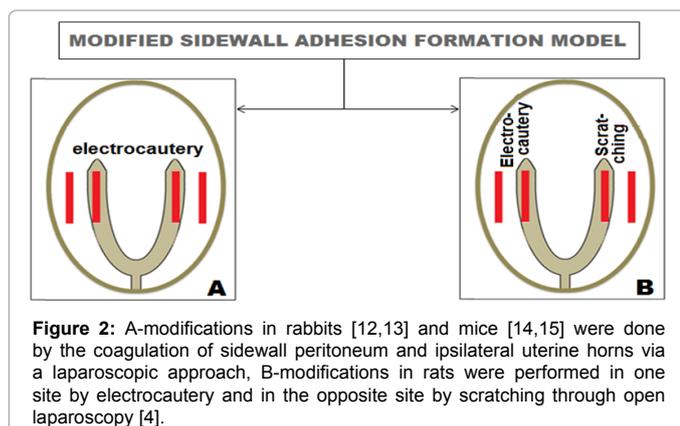


Figure 2: A-modifications in rabbits [12,13] and mice [14,15] were done by the coagulation of sidewall peritoneum and ipsilateral uterine horns via a laparoscopic approach, B-modifications in rats were performed in one site by electrocautery and in the opposite site by scratching through open laparoscopy [4].

Worldwide, experienced surgeons-practitioners and residents have also been involved in clinical trials of state-of-the art technologies and the evaluation of new surgical techniques. Key steps in the acquisition of clinically relevant results are the statement of questions and hypothesis building to set research goals as well as the elaboration of a study design.

As a prototype of our hypothesis building and testing tool we used a rabbit sidewall adhesion model [11]. Briefly, in their model Diamond et al. [11] performed a low midline laparotomy incision and exposed the uterine horns and abdominal/pelvic sidewalls. After several unsuccessful attempts with scraping or excisions of the peritoneum, and a combination of excisions of a 2 × 2-cm area of peritoneum and both layers of underlying muscle in conjunction with scraping of a contiguous 2-cm length of the uterine horn, the standardized postsurgical sidewall adhesion model was created by tacking together traumatized sidewall peritoneum and ipsilateral uterine horns using two interrupted sutures in each site (Figure 1). A piece of an absorbable adhesion barrier - Intercede (TC 7) was placed between these damaged lesions where opposite site was used own control. An evaluation of postsurgical adhesions was done after 2 weeks by scoring of adhesion extent, type and tenacity with successful improvements in sites with TC 7 application. Then this model and principle was used to evaluate many other adhesion barriers.

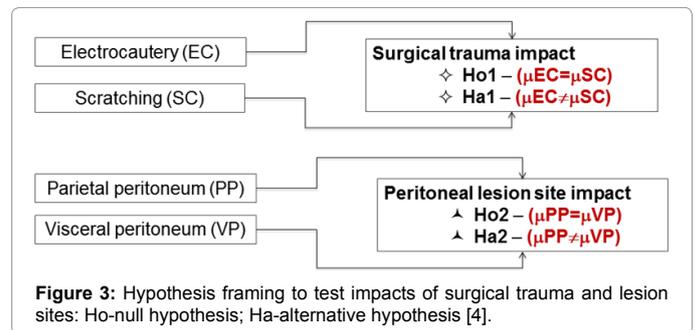


Figure 3: Hypothesis framing to test impacts of surgical trauma and lesion sites: Ho-null hypothesis; Ha-alternative hypothesis [4].

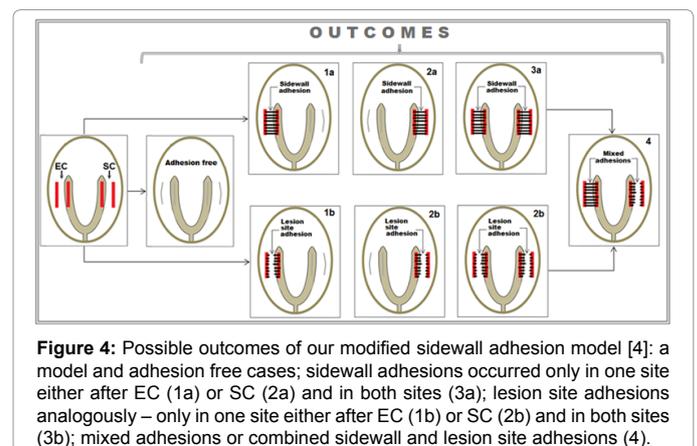


Figure 4: Possible outcomes of our modified sidewall adhesion model [4]: a model and adhesion free cases; sidewall adhesions occurred only in one site either after EC (1a) or SC (2a) and in both sites (3a); lesion site adhesions analogously – only in one site either after EC (1b) or SC (2b) and in both sites (3b); mixed adhesions or combined sidewall and lesion site adhesions (4).

Working hypotheses				Ho/Ha	Testing outcome	
Sidewall adhesions	Surgical trauma impact (EC vs SC)			Ho1	+ or -	
				Ha1		
Lesion site adhesions	Surgical trauma impact	EC-PP	vs	SC-PP	Ho2	+ or -
					Ha2	
		EC-VP	vs	SC-VP	Ho3	
				Ha3		
		EC-PP+VP	vs	SC-PP+VP	Ho4	+ or -
				Ha4		
Lesion site impact	PP-EC	vs	VP-EC	Ho5	+ or -	
						Ha5
	PP-SC	vs	VP-SC	Ho6	+ or -	
				Ha6		
PP-EC+SC	vs	VP-EC+SC	Ho7	+ or -		
			Ha7			

Notes: EC – electrocautery, SC-scratching, PP-parietal peritoneum, VP-visceral peritoneum (uterine horns), vs-versus.

Table 1: Types, framing and testing of working hypotheses.

This model has been modified to evaluate an impact of CO₂-pneumoperitoneum (Figure 2) by laparoscopic approach in rabbits [12,13] and in mice [14,15] at KULeuven and through open laparoscopy in rats [4] by our team at Moscow State University.

By application of one surgical damaging factor-electrocautery it was possible to test only an impact of CO₂-pneumoperitoneum [12-15]. Our model with two surgical factors such as electrocautery and scratching allowed us in addition to CO₂-pneumoperitoneum impact to test an impact of surgical trauma type and peritoneal lesion sites (Figure 3) with framing subsequent working hypotheses.

Our modified sidewall model increases our ability to make a comprehensively simultaneous analysis of an impact of several factors at the same experiment. An application of two different surgical trauma types in opposite sites in the same animal model allowed us to observe sidewall or lesion site adhesions occurring only in one site or both sites and a combination of these adhesions (Figure 4).

Subsequently, the types of working hypotheses, their framing and testing are presented in Table 1.

By evaluation of sidewall adhesions alone we could test an impact of type of surgical factor on postsurgical adhesion parameters. Additional evaluation of lesion site adhesions increased our ability to compare types of surgical factors such as dependence on lesion sites for instance, impact of surgical factor only on parietal peritoneum lesions or analogously on uterine horns (visceral peritoneum) or a combination of these factors. An impact of lesion sites was also studied in detail depending of the type of lesion sites by the same surgical factor and their combination.

In conclusion, based on our own experience and literature we recommend our modified model as a hypothesis building and testing tool in surgical experimental research.

Contributors

In design and description of this editorial opinion article all authors contributed equally.

References

1. Mynbaev OA, Tinelli A, Kosmas IP, Eliseeva MY, et al. The misleading adhesion formation model: a letter to the Editor. 2014 Jan 31.
2. Mynbaev OA, Rubl'oba KI, Liutova LV, Karabasova MA, Andreenko GV (1997) Role of local fibrinolytic activity of the uterine horns in rats in the pathogenesis of formation of postoperative adhesions. *Patol Fiziol Eksp Ter* 35-38.
3. Pismensky SV, Kalzhanov ZR, Eliseeva MY, Kosmas IP, Mynbaev OA (2011) Severe inflammatory reaction induced by peritoneal trauma is the key driving mechanism of postoperative adhesion formation. *BMC Surg* 11-30.
4. Mynbaev OA, Eliseeva MY, Kalzhanov ZR, Lyutova L, Pismensky SV, et al. (2013) Surgical trauma and CO₂-insufflation impact on adhesion formation in parietal and visceral peritoneal lesions. *Int J Clin Exp Med* 6: 153-165.
5. de Vries A, Marvik R, Kuhry E (2013) To perform operative procedures in an optimized local atmosphere: Can it reduce post-operative adhesion formation? *Int J Surg* 11: 1118-1122.
6. Ray NF, Denton WG, Thamer M, Henderson SC, Perry S (1998) Abdominal adhesiolysis: inpatient care and expenditures in the United States in 1994. *J Am Coll Surg* 186: 1-9.
7. Parker MC, Ellis H, Moran BJ, Thompson JN, Wilson MS, et al. (2001) Postoperative adhesions: ten-year follow-up of 12,584 patients undergoing lower abdominal surgery. *Dis Colon Rectum* 44: 822-829.
8. Tingstedt B, Isaksson J, Andersson R (2007) Long-term follow-up and cost analysis following surgery for small bowel obstruction caused by intra-abdominal adhesions. *Br J Surg* 94: 743-748.
9. Parker MC, Wilson MS, van Goor H, Moran BJ, Jeekel J, et al. (2007) Adhesions and colorectal surgery - call for action. *Colorectal Dis* 2: 66-72.
10. Mynbaev OA, Shi Z, Eliseeva MY, Medvedev MV, Andrea Tinelli (2013) A hypothesis building and testing model in surgical experimental researches. Minimally Invasive Surgery Week 2013 Annual Meeting & Endo Expo by SLS • AIMIS • CJMIS • CRSA • IPPS • ISGE • NESA • RAMSES • SMIT, at Reston VA (Washington DC, USA).
11. Diamond MP, Linsky CB, Cunningham T, Constantine B, diZerega GS, et al. (1987) A model for sidewall adhesions in the rabbit: reduction by an absorbable barrier. *Microsurgery* 8: 197-200.
12. Yesildaglar N, Koninckx PR (2000) Adhesion formation in intubated rabbits increases with high insufflation pressure during endoscopic surgery. *Hum Reprod* 15: 687-691.
13. Molinas CR, Koninckx PR (2000) Hypoxaemia induced by CO₂ or helium pneumoperitoneum is a co-factor in adhesion formation in rabbits. *Hum Reprod* 15: 1758-1763.
14. Yesildaglar N, Ordoñez JL, Laermans I, Koninckx PR (1999) The mouse as a model to study adhesion formation following endoscopic surgery: a preliminary report. *Hum Reprod* 14: 55-59.
15. Molinas CR, Mynbaev O, Pauwels A, Novak P, Koninckx PR (2001) Peritoneal mesothelial hypoxia during pneumoperitoneum is a cofactor in adhesion formation in a laparoscopic mouse model. *Fertil Steril* 76: 560-567.