

An experimental model of canine DNCB-induced allergic contact dermatitis: clinical and hematological features

Akram Zayerzadeh¹, Reza Avizeh^{2*}, Mohammad Khosravi³, Darioush Gharibi³ and
Mohammad Razijalali²

¹ DVSc Student of Small Animal Internal Medicine, Faculty of Veterinary Medicine, Shahid Chamran University of Ahvaz, Ahvaz, Iran

² Professor, Department of Clinical Sciences, Faculty of Veterinary Medicine, Shahid Chamran University of Ahvaz, Ahvaz, Iran

³ Associate Professor, Department of Pathobiology, Faculty of Veterinary Medicine, Shahid Chamran University of Ahvaz, Ahvaz, Iran

Received: 14.03.2022

Accepted: 11.06.2022

Abstract

Allergic contact dermatitis is an inflammatory skin reaction caused by direct contact with an offending substance. The aim of this study was to establish a suitable method for induction of allergic contact dermatitis in dogs for future studies. For this purpose, 1% and 2% dinitrochlorobenzene (DNCB) were injected with and without ethanol and olive oil or acetone and olive oil subcutaneously in the back of 8 BALB/c mice (4 equal groups). Then, based on the types and severity of the symptoms, it was decided to sensitize the dogs with only 2% DNCB. Finally, in two stages, 2% DNCB with or without dimethyl sulfoxide (DMSO) was injected with ethanol or acetone and olive oil subcutaneously in thoracic and scapular regions of five dogs. Two percent DNCB challenge caused clinical findings including erythema, edema and skin scaling, as well as pruritus and scratching behavior in the mice. Clinical findings in dogs developed in mild severity including redness and swelling within a few days. There was a significant increase in total blood leukocytes, neutrophils, lymphocytes, eosinophils and monocytes counts at seven days in dogs. Also statistical analysis showed that one week after DNCB injection, IgE expression increased significantly and this increase continued until day twenty-one. Based on the result of this study, intradermal injection of 100 µL of 2% DNCB mixture with DMSO (dissolved in a 4:1 mixture of ethanol and olive oil) was a suitable method for induction of allergic contact dermatitis in dogs. Furthermore, the scapular region was also a convenient location to prevent self-induced lesions.

Key words: Allergic contact dermatitis, Atopic dermatitis-like, Dinitrochlorobenzene, Dimethyl sulfoxide, Dog

* **Corresponding Author:** Reza Avizeh, Professor, Department of Clinical Sciences, Faculty of Veterinary Medicine, Shahid Chamran University of Ahvaz, Ahvaz, Iran
E-mail: avizeh@scu.ac.ir



© 2020 by the authors. Licensee SCU, Ahvaz, Iran. This article is an open access article distributed under the terms and conditions of the Creative Commons Attribution-NonCommercial 4.0 International (CC BY-NC 4.0 license) (<http://creativecommons.org/licenses/by-nc/4.0/>).

References

- Brinkhof, B., Spee, B., Rothuizen, J. & Penning, L. C. (2006). Development and evaluation of canine reference genes for accurate quantification of gene expression. *Analytical Biochemistry*, 356(1): 36–43.
- Cho, K. H., Parveen, A., Kang, M. C., Subedi, L., Lee, J. H., Park, S. Y., Jin, M. R., Yoon, H., Son, Y. K. & Kim, S. Y. (2018). *Pyrus ussuriensis* Maxim. leaves extract ameliorates DNCB-induced atopic dermatitis-like symptoms in NC/Nga mice. *Phytomedicine*, 48: 76-83.
- Deo, S. S., Mistry, K. J., Kakade, A. M. & Niphadkar, P. V. (2010). Role played by Th2 type cytokines in IgE mediated allergy and asthma. *Lung India*, 27(2): 66-71.
- Fang, L. W., Cheng, C. C., Hwang, T. S., Huang, W. C., Liou, C. J., Chen, W. C. & Wu, S. J. (2015). Danggui buxue tang inhibits 2,4-dinitrochlorobenzene: induced atopic dermatitis in mice. *Evidence-Based Complementary and Alternative Medicine*, 672891.
- Galvao, J., Davis, B., Tilley, M., Normando, E., Duchon, M. R., & Cordeiro, M. F. (2014). Unexpected low-dose toxicity of the universal solvent DMSO. *The FASEB Journal*, 28(3): 1317-1330
- Heo, J. Y., Cho, Y. S. & Cheon, H. G. (2010). Topical effects of roflumilast on 1-chloro-2,4-dinitrobenzene-induced atopic dermatitis-like skin lesions in NC/Nga mice. *Pharmazie*, 65(12): 906–912.
- Hwang, D. H., Koh, P. O., Kang, C. & Kim, E. (2021). *Rosa davurica* Pall. improves DNCB-induced atopic dermatitis in mice and regulated TNF- α /IFN- γ -induced skin inflammatory responses in HaCaT cells. *Phytomedicine*, 91: 153708.
- Jegal, J., Park, N. J., Bong, S. K., Jegal, H., Kim, S. N. & Yang, M. H., (2017). *Dioscorea quinqueloba* ameliorates Oxazolone- and 2,4-Dinitrochlorobenzene-induced Atopic Dermatitis Symptoms in murine models. *Nutrients*, 9(12): 1324.
- Jones S. E. & Versalovic J. (2009) Probiotic *Lactobacillus reuteri* biofilms produce antimicrobial and anti-inflammatory factors. *BMC Microbiology*, 9: 35.
- Kim, H. G., Kim K. S. & Oh, T. H. (2012). Anti-inflammatory effects of poly- γ -glutamic acid on DNCB-induced allergic contact dermatitis in dogs. *Journal of Veterinary Clinics*, 29(4): 283-296.
- Kimura, T. (1994). Dinitrochlorobenzene (DNCB)-induced delayed-type hypersensitivity reaction in the dorsal skin of hairless dogs. *Journal of Toxicologic Pathology*, 7: 229-235.
- Krawiec, D. R., & Gaafar, S. M. (1975). A comparative study of allergic and primary irritant contact dermatitis with dinitrochlorobenzene (DNCB) in dogs. *The Journal of Investigative Dermatology*, 65: 248- 251.
- Ku, J. M., Hong, S. H., Kim, S. R., Choi, H. S., Kim, H. I., Kim, D. U., Oh, S. M., Seo, H. S., Kim, T. Y., Shin, Y. C., Cheon, C. & Ko, S. G. (2018). The prevention of 2,4-dinitrochlorobenzene-induced inflammation in atopic dermatitis-like skin lesions in BALB/c mice by Jawoongo. *BMC Complementary and Alternative Medicine*, 18(1): 215.
- Lee, Y. K., Puong, K. Y., Ouwehand, A. C. & Salminen, S. (2003) Displacement of bacterial pathogens from mucus and Caco-2 cell surface by lactobacilli. *Journal of Medical Microbiology*, 52 (Pt 10): 925–930.
- Li, Y., Yu, H., Jin, Y., Li, M. & Qu, C. (2018). Verbascoside Alleviates Atopic Dermatitis-Like Symptoms in Mice via Its Potent Anti-Inflammatory Effect. *International Archives of Allergy and Immunology*, 175(4): 220-230.
- Miller, W. H., Griffin, C. E. & Campbell, K. L. (2013). *Muller and Kirk's Small Animal Dermatology* (7th Edition) Elsevier, Mosby, St. Louis, Missouri, USA. Pp 663-665.
- Mori, T., Tanimoto, Y., Ota, M., Masakado, T., Kitamoto, S., Saito, K., Isobe, N. & Kaneko, H. (2012). Comparison of cytokine profiles in bronchoalveolar lavage fluid of mice exposed to respiratory and contact sensitizers. *Journal of Toxicological Sciences*, 37(2): 337-343.
- Pagnini, C., Saeed, R., Bamias, G., Arseneau, K. O., Pizarro, T. T. & Cominelli, F. (2010) Probiotics promote gut health through stimulation of epithelial innate immunity. *Proceedings of the National Academy of Sciences of the United States of America*, 107: 454–459.
- Schlotter, Y. M., Rutten, V. P., Riemers, F. M., Knol, E. F. & Willemse, T. (2011). Lesional skin in atopic dogs shows a mixed Type-1 and Type-2 immune responsiveness. *Veterinary Immunology and Immunopathology*, 143(1-2): 20–26.

- Schmitz, S & Suchodolski, J. (2016). Understanding the canine intestinal microbiota and its modification by pro-, pre- and synbiotics – what is the evidence? *Veterinary Medicine and Science*, 2(2): 71–94.
- Schmitz, S., Garden, O. A., Werling, D. & Allenspach, K. (2012). Gene expression of selected signature cytokines of T cell subsets in duodenal tissues of dogs with and without inflammatory bowel disease. *Veterinary Immunology and Immunopathology*, 146(1): 87–91.
- Soo, I., Madsen, K. L., Tejpar, Q., Sydora, B. C., Sherbaniuk, R., Cinque, B. Marzio, L. Di., Grazia Cifone, M., Desimone, C. & Fedorak, R. N. (2008). VSL#3 probiotic upregulates intestinal mucosal alkaline sphingomyelinase and reduces inflammation. *Canadian Journal of Gastroenterology*, 22: 237–242.
- Wu, P. C., Chuo, W. H., Lin, S. C., Lehman, C. W., Lien, C. Z., Wu, C. S & Lin, C. C. (2019). Sclareol attenuates the development of atopic dermatitis induced by 2,4-dinitrochlorobenzene in mice. *Immunopharmacology and Immunotoxicology*, 41(1): 109-116.
- Yousif, A. D. & Abu-Raghif, A. R. (2020). The effect of topical dapsone in comparison with tacrolimus on DNCB induced atopic dermatitis in mice. *International Journal of Research in Pharmaceutical Sciences*, 11 (SPL4): 2050-2062.