

GLOBAL JOURNAL OF MEDICAL RESEARCH: A NEUROLOGY & NERVOUS SYSTEM Volume 23 Issue 1 Version 1.0 Year 2023 Type: Double Blind Peer Reviewed International Research Journal Publisher: Global Journals Online ISSN: 2249-4618 & Print ISSN: 0975-5888

Mexican Screening Test for Olfactory Dysfunction using Essential Oils

By Taniyama Natsuko, Carrillo-Mora Paul, Escobar-Barrios Marisa Selene, Hernández Rosales Paola, Mandujano-González Alberto, Ofelia Natsuko Taniyama López, Paul Carrillo Mora, Marisa Selene Escobar Barrios, Paola Hernández Rosales & Alberto Mandujano González

Abstract- Introduction: Olfaction is important for us to relate to the environment. Approximately 25% of the population over 40 years of age has an olfactory disorder. Olfactory alterations have a negative impact on quality of life, affecting self-esteem, generating depression, social isolation, and even altering eating habits. There are multiple etiologies: post-viral, traumatic, metabolic, secondary to medication, smoking and alcoholism, neurodegenerative diseases, among others. The tests available to screen for hyposmia include pens and books that give off scents, are expensive, difficult to access, require trained personnel to apply them and have been standardized in populations culturally different from the Mexican population, which makes it difficult to detect patients.

Objectives: To create, standardize and validate a screening test for olfactory alterations based on essential oils for the Mexican population.

Keywords: olfactory disorders, olfactory loss, screening, diagnosis.

GJMR-A Classification: DDC Code: 616.80471 LCC Code: RC347

MEX I CANSCREEN IN GTESTFORD LFACTORY DY SFUNCTIONUS IN GESSEN TIALOILS

Strictly as per the compliance and regulations of:



© 2023. Taniyama Natsuko, Carrillo-Mora Paul, Escobar-Barrios Marisa Selene, Hernández Rosales Paola, Mandujano-González Alberto, Ofelia Natsuko Taniyama López, Paul Carrillo Mora, Marisa Selene Escobar Barrios, Paola Hernández Rosales & Alberto Mandujano González. This research/review article is distributed under the terms of the Attribution-NonCommercial-NoDerivatives 4.0 International (CC BY-NC-ND 4.0). You must give appropriate credit to authors and reference this article if parts of the article are reproduced in any manner. Applicable licensing terms are at https://creativecommons.org/licenses/by-nc-nd/4.0/.

Mexican Screening Test for Olfactory **Dysfunction using Essential Oils**

Taniyama Natsuko ^α, Carrillo-Mora Paul ^α, Escobar-Barrios Marisa Selene ^ρ, Hernández Rosales Paola ^ω, Mandujano-González Alberto[¥], Ofelia Natsuko Taniyama López[§], Paul Carrillo Mora^x, Marisa Selene Escobar Barrios^{*}, Paola Hernández Rosales[®] & Alberto Mandujano González^{^ζ}

Abstract- Introduction: Olfaction is important for us to relate to the environment. Approximately 25% of the population over 40 years of age has an olfactory disorder. Olfactory alterations have a negative impact on guality of life, affecting self-esteem, generating depression, social isolation, and even altering eating habits. There are multiple etiologies: post-viral, traumatic, metabolic, secondary to medication, smoking and alcoholism, neurodegenerative diseases, among others. The tests available to screen for hyposmia include pens and books that give off scents, are expensive, difficult to access, require trained personnel to apply them and have been standardized in populations culturally different from the Mexican population, which makes it difficult to detect patients.

Objectives: To create, standardize and validate a screening test for olfactory alterations based on essential oils for the Mexican population.

Methods: A smell test was created with the aromas: lemon, cinnamon, chocolate, coffee and mint, and it was standardized in a previous work, with the application of 630 tests. Subsequently, the results of application were compared with the results obtained by applying the University of Pennsylvania Smell Identification Test and the identification part of the Sniffin Sticks test.

Results: The test obtained a sensitivity of 0.93, specificity of 0.77, negative predictive value of 0.9 and positive of 0.77, data compared with the Gold Standard test, demonstrating its noninferiority.

Conclusion: The screening test with essential oils is a valid, reliable, affordable, fast and easy to apply test to detect olfactory alterations in the Mexican population.

Keywords: olfactory disorders, olfactory loss, screening, diagnosis.

INTRODUCTION I.

he ability to smell allows us to enjoy food, relate to others and even protect ourselves from danger. Moreover, because of its connections with the limbic system, it allows us to evoke intense emotions and memories¹.

e-mail: natsukotaniyama@gmail.com

Olfactory disorders are classified into qualitative problems (dysosmias, i.e., altered olfactory perception) and quantitative problems (where the intensity of olfactory perception is affected and includes hyposmia and anosmia)².

Of people over 40 years of age, 25% have some olfactory disorder, which increases with age, reaching a prevalence of 40-62% in people over 80 years of age³,4. The causes of these conditions are: nasosinusal alterations. post-infectious olfactory dysfunction, presbymosmia. post-traumatic. secondarv to smoking alcoholism, medication, chronic and neurodegenerative diseases, metabolic and genetic diseases^{5,6}.

The presence of olfactory alterations has an impact on the quality of life of sufferers, generating depression, social isolation, feelings of vulnerability, selfesteem problems and causing insecurity, since up to 60% of those affected report difficulty in noticing a gas leak or the presence of smoke. Eating habits, personal hygiene and sexual performance are affected^{7,8,9.}

Olfactory impairment is a predictor of mortality, since subjects > 60 years of age with anosmia are 3 times more likely to die at 5 years than normosmic patients¹⁰.

Existing tests for screening for hyposmia are variable, including pens such as the Sniffin Sticks (SS) test^{11,12}, booklets such as the University of Pennsylvania Smell Identification Test (UPSIT) or disks that give off scents^{13,14}. They are expensive, not very accessible in Mexico and require training to perform them. These tests have been standardized in populations culturally very different from the Mexican population, so the type of aromas 15,16,17 their familiarity and therefore their identification are different in Mexico. This makes it difficult to detect patients and to know the prevalence of hyposmia in our population.

So far no studies have been conducted on the usefulness of using commercial essential oils in the detection of hyposmia, which are readily available, inexpensive, and easy and quick to use.

In Mexico there is no standardized test for the screening of olfactory alterations, so this work proposes the development and validation of a test of essential oils for this purpose.

Author $\alpha \ \rho \ \omega \ \neq \ v \ \theta \ \zeta$: Otolaryngology Head and Neck Surgery Department, National Rehabilitation Institute "Luis Guillermo Ibarra Ibarra", Mexico City, Mexico.

Author $\sigma \chi$: Neuroscience Division, National Rehabilitation Institute "Luis Guillermo Ibarra Ibarra", Mexico City, Mexico.

Corresponding Author §: Ofelia Natsuko Taniyama-López, Otolaryngology Head and Neck Surgery Department, National Rehabilitation Institute "Luis Guillermo Ibarra Ibarra". Calzada México-Xochimilco 289, Tlalpan, México City, Mexico, 14389.

II. MATERIALS AND METHODS

A descriptive study was carried out, in which two groups were created, one with subjective hyposmia of different etiologies and another age and gender matched control group made up of healthy people who denied symptomatic hyposmia.

The patients were recruited at the National Rehabilitation Institute-LGII within the period June 2021-March 2022.

The case group included subjects older than 18 years with hyposmia caused by nasal conditions: attic septal deviation, acute sinusitis, chronic sinusitis with and without polyps, nasal tumors, upper airway infections including COVID 19 corroborated by PCR, Parkinson's disease.

All subjects underwent nasal endoscopy to corroborate the presence of nasal conditions. Subjects with active smoking and those with a history of nose and sinus surgery were excluded.

The following scales were applied to each case with olfactory alterations: Sinonasal Outcome 22 (SNOT 22) ^{19,20,21} and Questionnaire of Olfactory Disorders-Negative Statements (QOD NS), ^{22,23}and an endoscopic scale was used to evaluate the state of the olfactory cleft: The olfactory cleft endoscopy scale (OCES).²⁴

To evaluate the olfactory function, the University of Pennsylvania Smell Identification Test (UPSIT) and the Sniffin Sticks test, the identification subset with 16 aromas, were applied. In this same consultation, the test with essential oils was applied to each participant. The order of the olfactory evaluation was as follows: UPSIT, essential oil test, Sniffin Sticks.

Between each test a 5-minute break was taken to prevent olfactory memory from influencing the results and to allow for mental relaxation of the participants.

In a previous investigation, this test of commercial essential oils was standardized²⁵. This test consists briefly in the use of 5 aromas: lemon, chocolate, cinnamon, coffee, mint and a control with no odor (Image 1).

The interpretation of this test will be only if the patient correctly or incorrectly identified each aroma, therefore it fluctuates between 0-5, if the subject obtains 0=anosmia, 1,2,3= hyposmia and 4,5 points= normosmia (Image 2). (Image 2).

The statistical package Prisma was used. Statistical significance was considered with a p < 0.05.

For olfactory function, each participant obtained three scores derived from the tests used: number of essential oils correctly identified, score on the University of Pennsylvania Smell Identification Test and score on the Sniffin Sticks test and also obtained an olfactory diagnosis from each test: Normosmia, hyposmia or anosmia.

Fisher's exact test was used to compare the numbers of anosmic, hyposmic and normosmic

subjects with scores of 0- 5 correctly identified oils, these results were compared with the UPSIT and Sniffin Sticks scores.

Sensitivity and specificity of test results were determined by performing 2 x 2 contingency tables.

III. Results

A total of 33 subjects were analyzed, 17 in the case group and 16 in the control group. The case group consisted of 7 women and 10 men, the age range was 18-82 years with a mean age of 41 years. The control group consisted of 7 women and 9 men, the age range was 18-81 years with a mean age of 46 years.

Regarding the cause of hyposmia in the case group, 2 had no identifiable cause, 5 were due to COVID 19, 3 due to nasosinus conditions, 4 due to Parkinson's disease and 3 due to cranioencephalic trauma.

The presence of nasal alterations, tumors or infectious diseases was ruled out in all the controls by nasal endoscopy. The mean score on the endoscopic evaluation scale of the olfactory sulcus "OCES", in the cases, was 1.1 points in the right nostril and 1.2 in the left nostril.

Regarding quality of life, in the group of cases, the range of scores was 8-33 points, with a mean of 25, which in percentage of affection of 1-100 translates into 14%-57.8%.

For the Sniffin Sticks test the scores were 1 - 15 in the case group and for the controls 11- 15.

In the UPSIT test the score in the case group was 9-29 and in the controls from 25 to 40. With this test in the group of cases all had some degree of hyposmia and in the group of controls 4 had mild microsmia, 2 with moderate and 1 with severe, the subjects with mild and moderate microsmia had no alterations in the other two tests and the one with severe hyposmia was normal in the Sniffin Sticks test and with hyposmia in the essential oils test.

For the essential oils test, a score of 4 and 5 points was taken as normal, hyposmia 1-3 and anosmia 0. An additional point was added to be taken into account in the final score; for each aroma, the intensity with which each aroma was perceived was questioned and if the subject did not perceive an aroma or if 3 of the aromas were perceived with slight intensity, one point was subtracted from the total number of correctly identified aromas (this variant was called modified oils test in the analysis).

Table 1 shows the olfactory results obtained with the different tests, sensitivity, specificity, positive and negative predictive value and confidence interval calculated with Fisher's test.

IV. DISCUSSION

The screening test for olfactory disorders with essential oils proved to be an option with acceptable

sensitivity and specificity. Figures ranging from 0.88-0.93 and 0.41-0.77 respectively.

In 2018 in Barcelona Campabadal et al. determined the sensitivity and specificity of the UPSIT test Spanish version in a population with Parkinson's disease: 97 subjects, and healthy controls: 65 and determined that the sensitivity was 81.4% and specificity 100% with a cut-off point \leq 25 points.²⁶

Hummel and his team in 1997 created the Sniffin Sticks test, applied it to a group of 104 healthy subjects (52 women, 52 men, mean age 49.5 years, range 18-84 years) and compared it to an established measure of olfactory performance; the Connecticut Clinical Chemosensory Research Center Test, CCCRC²⁷. The use of the different subsets separately has been found to have a sensitivity and specificity of 84%.²⁸

Sorokowska et al. in 2019 conducted a multicenter study in Germany with 333 subjects with olfactory disturbances of different etiologies, aged 12-88 years. In whom they evaluated the clinical utility of employing a test created by the researchers "Q-Sticks test", a test composed of the aromas of clove, coffee and roses. Their test obtained a sensitivity of 91.8% and a specificity of 92%²⁹.

A retrospective study was carried out in Germany in 2016 with 613 subjects with an age range of 18-96 years, they included subjects with olfactory disturbances (464) of different etiologies and controls (149), to whom they applied the Sniffin Sticks olfactory identification subset containing 16 scents. All participants underwent nasal endoscopy and medical history. They created a score for each aroma based on the following division: % of subjects with normosmia who correctly identified it between % of subjects with identifiable cause of hyposmia who correctly identified it and called it "odor specificity score", then using a calculated ABC analysis which is a classification method used in economics, which allows identifying items that have an important impact on an overall value they selected 3 aromas. Cinnamon, fish and banana were correctly identified by the largest number of normosmic subjects, with this battery of tests they obtained a sensitivity of 80.4%, specificity of 84.3% and a negative predictive value of 91.3%³⁰.

It is important to mention that, during the analysis of the test results, we noticed the differences between the tests at the time of interpreting their results, i.e., the UPSIT test graduates the level of hyposmia into mild microsmia, moderate or severe hyposmia, while the Sniffin Sticks and the essential oils test only identifies normosmia, hyposmia or anosmia. Because of these differences, subjects who obtained normosmia in the last two tests, but mild microsmia in the UPSIT test were considered to have normosmia, thus obtaining different figures for sensitivity, specificity, negative predictive value and positive predictive value. In addition, when taking into account the intensity of aroma perception in the oil test; "modified oil test" increased all the parameters analyzed.

The essential oil screening test we propose is an effective method, easily accessible, cost-effective and quick to apply. With respect to its sensitivity and specificity, it is very similar to that shown by other tests, thus determining its non-inferiority.

The great limitation of this study is the low number of samples; however, due to the favorable behavior of the data, it can be inferred that by increasing the number of subjects, similar results will be obtained.

V. Conclusion

This is a first step in the detection of olfactory disorders in Mexico; however, future research is needed to extend the level of diagnosis, in order to obtain a test with which to follow up patients or even determine the efficacy of certain treatments.

References Références Referencias

- Croy, I., Nordin, S., & Hummel, T. (2014). Olfactory disorders and quality of life--an updated review. Chemical Senses, 39(3), 185–194. https://doi.org/ 10.1093/CHEMSE/BJT072
- Boesveldt, S., Postma, E. M., Boak, D., Welge-Luessen, A., Schöpf, V., Mainland, J. D., Martens, J., Ngai, J., & Duffy, V. B. (2017). Anosmia-A Clinical Review. Chemical Senses, 42(7), 513–523. https://doi.org/10.1093/CHEMSE/BJX025
- Murphy, C., Schubert, C. R., Cruickshanks, K. J., Klein, B. E. K., Klein, R., & Nondahl, D. M. (2002). Prevalence of olfactory impairment in older adults. JAMA, 288(18), 2307–2312. https://doi.org/10.1001/ JAMA.288.18.2307
- Liu, G., Zong, G., Doty, R. L., & Sun, Q. (2013). Prevalence and risk factors of taste and smell impairment in a nationwide representative sample of the US population: a cross-sectional study. National Health and Nutrition Examination Survey, 2013– 2014. https://doi.org/10.1136/bmjopen-2016
- 5. Devere, R. (2017). Disorders of Taste and Smell. Continuum (Minneapolis, Minn.), 23(2, Selected Topics in Outpatient Neurology), 421–446. https://doi.org/10.1212/CON.000000000000463
- Imam, S. A., Lao, W. P., Reddy, P., Nguyen, S. A., & Schlosser, R. J. (2020). Is SARS-CoV-2 (COVID-19) postviral olfactory dysfunction (PVOD) different from other PVOD? World Journal of Otorhinolaryngology -Head and Neck Surgery, 6(Suppl 1), S26–S32. https://doi.org/10.1016/J.WJORL.2020.05.004
- Miwa, T., Furukawa, M., Tsuhatani, T., Costanzo, R. M., DiNardo, L. J., & Reiter, E. R. (2001). Impact of olfactory impairment on quality of life and disability. Archives of Otolaryngology--Head & Neck Surgery,

127(5), 497–503. https://doi.org/10.1001/ ARCHOTOL.127.5.497

- Liu, G., Zong, G., Doty, R. L., & Sun, Q. (2013). Prevalence and risk factors of taste and smell impairment in a nationwide representative sample of the US population: a cross-sectional study. National Health and Nutrition Examination Survey, 2013– 2014. https://doi.org/10.1136/bmjopen-2016
- Hummel, T., & Nordin, S. (2005). Olfactory disorders and their consequences for quality of life. In Acta Oto-Laryngologica (Vol. 125, Issue 2, pp. 116–121). https://doi.org/10.1080/00016480410022787
- Pinto, J. M., Wroblewski, K. E., Kern, D. W., Schumm, L. P., & McClintock, M. K. (2014).
 Olfactory Dysfunction Predicts 5-Year Mortality in Older Adults. PLOS ONE, 9(10), e107541. https://doi.org/10.1371/JOURNAL.PONE.0107541
- Hummel, T., & Welge-Lüessen, A. (2006). Assessment of Olfactory Function. In Taste and Smell (1st ed., Vol. 63, pp. 84–98). KARGER. https://doi.org/10.1159/000093752)
- Rumeau, C., Nguyen, D. T., & Jankowski, R. (2016). How to assess olfactory performance with the Sniffin' Sticks test(®). European Annals of Otorhinolaryngology, Head and Neck Diseases, 133(3), 203–206. https://doi.org/10.1016/J.ANORL. 2015.08.004
- 13. Doty, R. L., Shaman, P., & Dann, M. (1984). Development of the University of Pennsylvania Smell Identification Test: standardized а microencapsulated test of olfactory function. Physiology Behavior, 32(3). & 489-502. https://doi.org/10.1016/0031-9384(84)90269-5
- 14. Doty R.L. The Smell identification testTM administration manual 3rd Ed. University of Pennsylvania. 1995.
- Lötsch, J., Ultsch, A., & Hummel, T. (2016). How many and which odor identification items are needed to establish normal olfactory function? Chemical Senses, 41(4), 339–344. https://doi.org/ 10.1093/chemse/bjw006
- Yáñez, C., Mora, N., & Nurko, B. (2004). A B C Prueba corta de olfato a utilizarse como una prueba de diagnóstico confiable. 49, 82–86.
- Yanagimachi, M., Takahashi, I., Bernier, F., Takahashi, K., Kurauchi, S., Mikuniya, Y., Sasaki, A., Komine, T., Matsubara, A., & Nakaji, S. (2017). Olfactory function evaluation in a 1102 communitydwelling 20-90-year old Japanese population in relation with age, sex and mental decline. Alzheimer's, Dementia & Cognitive Neurology, 1(3). https://doi.org/10.15761/ADCN.1000117
- Hummel, T., & Nordin, S. (2005). Olfactory disorders and their consequences for quality of life. In Acta Oto-Laryngologica (Vol. 125, Issue 2, pp. 116–121). https://doi.org/10.1080/00016480410022787

- Piccirillo, J. F., Merritt, M. G., & Richards, M. L. (2002). Psychometric and clinimetric validity of the 20-Item Sino-Nasal Outcome Test (SNOT-20). Otolaryngology--Head and Neck Surgery: Official Journal of American Academy of Otolaryngology-Head and Neck Surgery, 126(1), 41–47. https://doi.org/10.1067/MHN.2002.121022
- Hopkins, C., Gillett, S., Slack, R., Lund, V. J., & Browne, J. P. (2009). Psychometric validity of the 22-item Sinonasal Outcome Test. Clinical Otolaryngology : Official Journal of ENT-UK ; Official Journal of Netherlands Society for Oto-Rhino-Laryngology & Cervico-Facial Surgery, 34(5), 447– 454. https://doi.org/10.1111/J.1749-4486.2009. 01995.X
- 21. de los Santos, G., Reyes, P., del Castillo, R., Fragola, C., & Royuela, A. (2015). Cross-cultural adaptation and validation of the sino-nasal outcome test (SNOT-22) for Spanish-speaking patients. European Archives of Oto-Rhino-Laryngology: Official Journal of the European Federation of Oto-Rhino-Laryngological Societies (EUFOS): Affiliated with the German Society for Oto-Rhino-Laryngology - Head and Neck Surgery, 272(11), 3335–3340. https://doi.org/10.1007/S00405-014-3437-0
- Frasnelli, J., & Hummel, T. (2005). Olfactory dysfunction and daily life. European Archives of Oto-Rhino-Laryngology : Official Journal of the European Federation of Oto-Rhino-Laryngological Societies (EUFOS) : Affiliated with the German Society for Oto-Rhino-Laryngology - Head and Neck Surgery, 262(3), 231–235. https://doi.org/10.1007/S00405-004-0796-Y
- Mattos, J. L., Schlosser, R. J., Storck, K. A., & Soler, Z. M. (2017). Understanding the relationship between olfactory-specific quality of life, objective olfactory loss, and patient factors in chronic rhinosinusitis. International Forum of Allergy and Rhinology, 7(7), 734–740. https://doi.org/10.1002/ alr.21940
- Schlosser, R. J., Smith, T. L., Mace, J. C., Alt, J. A., Beswick, D. M., Mattos, J. L., Ramakrishnan, V., Massey, C., & Soler, Z. M. (2021). The Olfactory Cleft Endoscopy Scale: a multi-institutional validation study in chronic rhinosinusitis. Rhinology, 59(2), 181–190. https://doi.org/10.4193/RHIN20.307
- 25. Natsuko Taniyama-López, O., Carrillo-Mora, P., Selene Escobar-Barrios, M., Hernández-Rosales, P., Mandujano-González, A., & Ofelia Natsuko Taniyama López, C. (2022). Estandarización de una prueba de tamizaje de alteraciones del olfato con aceites esenciales Standardization of a screening test of olfactory alterations with essential oils. https://doi.org/10.24245/aorl.v67i1.7502
- 26. Campabadal, A., Segura, B., Baggio, H. C., Abos, A., Uribe, C., Garcia-Diaz, A. I., Marti, M. J., Valldeoriola, F., Compta, Y., Bargallo, N., & Junque,

C. (2019). Diagnostic Accuracy, Item Analysis and Age Effects of the UPSIT Spanish Version in Parkinson's Disease. Archives of Clinical Neuropsychology: The Official Journal of the National Academy of Neuropsychologists, 34(4), 435–444. https://doi.org/10.1093/ARCLIN/ACY053

- 27. Hummel, T., Sekinger, B., Wolf, S. R., Pauli, E., & Kobal, G. (1997). "Sniffin" sticks': olfactory performance assessed by the combined testing of odor identification, odor discrimination and olfactory threshold. Chemical Senses, 22(1), 39–52. https://doi.org/10.1093/CHEMSE/22.1.39
- Lötsch, J., Reichmann, H., & Hummel, T. (2008). Different odor tests contribute differently to the evaluation of olfactory loss. Chemical Senses, 33(1), 17–21. https://doi.org/10.1093/CHEMSE/BJM058
- Sorokowska, A., Oleszkiewicz, A., Minovi, A., Konnerth, C. G., & Hummel, T. (2019). Fast screening of olfactory function using the q-sticks test. ORL, 81(5–6), 245–251. https://doi.org/ 10.1159/000500559
- Lötsch, J., Ultsch, A., & Hummel, T. (2016a). How many and which odor identification items are needed to establish normal olfactory function? Chemical Senses, 41(4), 339–344. https://doi.org/ 10.1093/chemse/bjw006