## **Scopus**

### Documents

Irnawati, I.<sup>a b</sup>, Windarsih, A.<sup>c</sup>, Putri, A.R.<sup>d</sup>, Fadzillah, N.A.<sup>e</sup>, Azmi, A.A.<sup>f</sup>, Kusbandari, A.<sup>g</sup>, Rohman, A.<sup>h</sup>

The use of spectroscopic methods in combination with multivariate data analysis for determination of omega fatty acids: A review

(2024) Journal of Applied Pharmaceutical Science, 14 (1), pp. 45-53.

DOI: 10.7324/JAPS.2024.139258

<sup>a</sup> Universitas Gadjah Mada, Yogyakarta, Indonesia

<sup>b</sup> Study Program of Pharmacy, Faculty of Pharmacy, Halu Oleo University, Kendari, Indonesia

<sup>c</sup> Research Center for Food Technology and Processing (PRTPP), National Research and Innovation Agency (BRIN), Yogyakarta, Indonesia

<sup>d</sup> Departement of Pharmacy, Faculty of Medicine, Universitas Brawijaya, Malang, Indonesia

<sup>e</sup> International Institute for Halal Research and Training (INHART), International Islamic University Malaysia, Kuala Lumpur, Malaysia

<sup>†</sup> Halal Products Research Institute, University Putra Malaysia, Serdang, Malaysia

<sup>g</sup> Division of Pharmaceutical Analysis and Medicinal Chemistry, Faculty of Pharmacy, Universitas Ahmad Dahlan, Yogyakarta, Indonesia

<sup>h</sup> Department of Pharmaceutical Chemistry, Faculty of Pharmacy, Gadjah Mada University, Yogyakarta, Indonesia

#### Abstract

Omega-3 fatty acids ( $\omega$ -3 FAs), typically found in fish oils and marine-based products, are important fatty acids due to their beneficial activities toward human health, such as anti- inflammation, immune-stimulant, lowering the risk of cardiovascular disease and reducing blood pressure. Therefore, the determination of  $\omega$ -3 FAs for quality control of products containing these FAs is very important. Molecular spectroscopic methods offered simple, fast, and reliable analytical methods for quality controls of food and pharmaceutical products containing  $\omega$ -3 FAs since a large amount of information could be retrieved from molecular spectra. This review highlighted the employment of molecular spectroscopy such as near-infrared (NIR), Fourier transform infrared (FTIR), Raman, and nuclear magnetic resonance (NMR) spectrometer combined with multivariate data analysis or chemometrics for analysis of  $\omega$ -3 FAs in fish oil-based products. From this review, it is reported that the combination of molecular spectroscopy and chemometrics could be used as effective analytical techniques for the analysis of  $\omega$ -3 FAs, especially eicosapentaenoic acid (C20:5,  $\omega$ -3) and docosahexaenoic acid (C22:6,  $\omega$ -3), with high accuracy and high precision. The results of quantitative analysis of  $\omega$ -3 FAs from NIR, FTIR, Raman, and NMR were comparable to those reference results obtained from gas chromatography-mass spectrometry measurement. In the future, collaborative studies through proficiency testing should be performed to get standardized methods based on molecular spectroscopy and chemometrics. ( $\omega$  2024 Irnawati Irnawati et al. This is an open access article distributed under the terms of the Creative Commons Attribution 4.0 International License (https://creativecommons.org/licenses/by/4.0/).

#### Author Keywords

chemometrics; DHA; EPA; Molecular spectroscopy; omega fatty acids

#### References

- Park, SE, Yu, HY, Ahn, S.
   Development and validation of a simple method to quantify contents of phospholipids in krill oil by Fourier-transform infrared spectroscopy (2022) *Foods*, 11, p. 41.
  - 1
- Pasini, F, Gómez-Caravaca, AM, Blasco, T, Cvejić, J, Caboni, MF, Verardo, V.
   Assessment of lipid quality in commercial omega-3 supplements sold in the French market

   (2022) Biomolecules, 12, p. 1361.

(2022) *Biomolecules*, 12, p. 1361. 2

Vongsvivut, J, Heraud, P, Gupta, A, Puri, M, McNaughton, D, Barrow, CJ.
 FTIR microspectroscopy for rapid screening and monitoring of polyunsaturated fatty acid production in commercially valuable marine yeasts and protists (2013) *Analyst*, 138, pp. 6016-6031.

- Innes, JK, Calder, PC.
   Marine omega-3 (N-3) fatty acids for cardiovascular health: an update for 2020 (2020) Int J Mol Sci, 21, pp. 1-21.
   4
- Mozaffari, H, Daneshzad, E, Larijani, B, Bellissimo, N, Azadbakht, L.
   Dietary intake of fish, n-3 polyunsaturated fatty acids, and risk of inflammatory bowel disease: a systematic review and meta-analysis of observational studies (2020) *Eur J Nutr*, 59, pp. 1-17.
- Giacobbe, J, Benoiton, B, Zunszain, P, Pariante, CM, Borsini, A.
   The anti-inflammatory role of omega-3 polyunsaturated fatty acids metabolites in pre-clinical models of psychiatric, neurodegenerative, and neurological disorders (2020) *Front Psychiatry*, 11, p. 122.
- Gao, F, Han, LJ, Liu, X.
   Vibration spectroscopic technique for species identification based on lipid characteristics

   (2017) Int J Agric Biol Eng, 10, pp. 255-268.
   7
- Alinafiah, SM, Azlan, A, Ismail, A, Rashid, NKMA.
   Method development and validation for omega-3 fatty acids (DHA and EPA) in fish using gas chromatography with flame ionization detection (GC-FID) (2021) *Molecules*, 26, p. 6592.
- Yi, T, Li, SM, Fan, JY, Fan, LL, Zhang, ZF, Luo, P
   Comparative analysis of EPA and DHA in fish oil nutritional capsules by GC-MS (2014) *Lipids Health Dis*, 13, pp. 1-6.
   9
- Kiełbasa, A, Buszewski, B, Gadzała-Kopciuch, R.
   A novel non-derivatization HPLC/UV method for the determination of some n-3 free fatty acids in breast milk matrix

   (2022) *Microchem J*, 181, p. 107789.
   10
- Viswanathan, S, Verma, PRP, Ganesan, M, Manivannan, J.
   A novel liquid chromatography/tandem mass spectrometry (LC–MS/MS) based bioanalytical method for quantification of ethyl esters of eicosapentaenoic acid (EPA) and docosahexaenoic acid (DHA) and its application in pharmacokinetic study

   (2017) J Pharm Biomed Anal, 141, pp. 250-261.
   11
- Rohman, A, Rawar, EA, Sudevi, S, Nurrulhidayah, AF, Windarsih, A.
   The use of chemometrics in combination with molecular spectroscopic and chromatographic methods for authentication of curcuma species: a review (2020) *Food Res*, 4, pp. 1850-1858.
   12
- Breuer, D.
   Molecular spectroscopy in the ultraviolet and visible range, pp. 1-5.
   13. Hoboken (NJ): John Wiley & Sons
- Rohman, A, Windarsih, A. The application of molecular spectroscopy in combination with chemometrics for

halal authentication analysis: a review (2020) *Int J Mol Sci*, 21, pp. 1-18. 14

- Pollo, BJ, Teixeira, CA, Belinato, JR, Furlan, MF, Cunha IC de, M, Vaz, CR Chemometrics, comprehensive two-dimensional gas chromatography and "omics" sciences: basic tools and recent applications (2021) *Trends Anal Chem*, 134, p. 116111.
- Moros, J, Garrigues, S, De Guardia, M.
   Vibrational spectroscopy provides a green tool for multi-component analysis (2010) *Trends Anal Chem*, 29, pp. 578-591.
   16
- Paul, A, De, P, Harrington, B.
   Chemometric applications in metabolomic studies using chromatography-mass spectrometry

   (2021) Trends Anal Chem, 135, p. 116165.
   17
- Lee, LC, Liong, CY, Jemain, AA.
   A contemporary review on data preprocessing (DP) practice strategy in ATR-FTIR spectrum
   (2017) Chemom Intell Lab Syst, 163, pp. 64-75.
   18
- Rinnan, Å, van den, Berg F, Engelsen, SB.
   Review of the most common pre-processing techniques for near-infrared spectra (2009) *Trends Anal Chem*, 28, pp. 1201-1222.
   19
- Rohman, A, Che Man, YB, Nurrulhidayah, AF.
   Fourier-transform infrared spectra combined with chemometrics and fatty acid composition for analysis of pumpkin seed oil blended into olive oil (2015) *Int J Food Prop*, 18, pp. 1086-1096.
   20
- Peris-Díaz, MD, Krężel, A.
   A guide to good practice in chemometric methods for vibrational spectroscopy, electrochemistry, and hyphenated mass spectrometry

   (2021) Trends Anal Chem, 135, p. 116157.
   21
- Rohman, A, Windarsih, A, Hossain, MAM, Johan, MR, Ali, ME, Fadzilah, NA.
   Application of near- and mid-infrared spectroscopy combined with chemometrics for discrimination and authentication of herbal products: a review (2019) *J Appl Pharm Sci*, 9, pp. 137-147.
   22
- Brereton, RG.
   Consequences of sample size, variable selection, and model validation and optimisation, for predicting classification ability from analytical data (2006) *Trends Anal Chem*, 25, pp. 1103-1111.
   23
- Saad, AS, Elzanfaly, ES, Halim, MK, Kelani, KM.
   Comparing the predictability of different chemometric models over UV-spectral data of isoxsuprine and its toxic photothermal degradation products (2019) Spectrochim Acta Part A Mol Biomol Spectrosc, 219, pp. 444-449.
   24

26

- Bekhit, MY, Grung, B, Mjøs, SA.
   Determination of omega-3 fatty acids in fish oil supplements using vibrational spectroscopy and chemometric methods
   (2014) *Appl Spectrosc*, 68, pp. 1190-1200.
   25
- Cascant, MM, Breil, C, Fabiano-Tixier, AS, Chemat, F, Garrigues, S, de la Guardia, M. Determination of fatty acids and lipid classes in salmon oil by near infrared spectroscopy (2018) Food Chem, 239, pp. 865-871.
- Karunathilaka, SR, Choi, SH, Mossoba, MM, Yakes, BJ, Brückner, L, Ellsworth, Z Rapid classification and quantification of marine oil omega-3 supplements using ATR- FTIR, FT-NIR and chemometrics (2019) *J Food Compos Anal*, 77, pp. 9-19.
- Van der Merwe, S, Manley, M, Wicht, M.
   Enhancing near infrared spectroscopy models to identify omega-3 fish oils used in the nutraceutical industry by means of calibration range extension (2018) *J Near Infrared Spectrosc*, 26, pp. 245-261.
- Lopes, TIB, Pereira, ES, dos Freitas, DS, Oliveira, SL, Alcantara, GB.
   Spectral profiles of commercial omega-3 supplements: an exploratory analysis by ATR-FTIR and 1H NMR
   (2020) *J Food Sci Technol*, 57, pp. 1251-1257.
   29
- Toyoda, K, Yamanoue, M, Ihara, I, Hu, X.
   ATR-FTIR evaluation of important fatty acid profile in Japanese black cattle beef (2012) 58th International Congress of Meat Science and Technology, 30. August 12–17; Montreal, Canada

 Hernández-Martínez, M, Gallardo-Velázquez, T, Osorio-Revilla, G, Almaraz-Abarca, N, Ponce- Mendoza, A, Vásquez-Murrieta, MS.
 Prediction of total fat, fatty acid composition and nutritional parameters in fish fillets using MID-FTIR spectroscopy and chemometrics (2013) *LWT Food Sci Technol*, 52, pp. 12-20.
 31

- Karunathilaka, SR, Yakes, BJ, Choi, SH, Brückner, L, Mossoba, MM.
   Comparison of the performance of partial least squares and support vector regressions for predicting fatty acids and fatty acid classes in marine oil dietary supplements by using vibrational spectroscopic data (2020) *J Food Prot*, 83, pp. 881-889.
   32
- Vongsvivut, J, Miller, MR, McNaughton, D, Heraud, P, Barrow, CJ.
   Rapid discrimination and determination of polyunsaturated fatty acid composition in marine oils by FTIR spectroscopy and multivariate data analysis (2014) Food Bioprocess Technol, 7, pp. 2410-2422.
   33
- Daoud, S, Bou-maroun, E, Dujourdy, L, Waschatko, G, Billecke, N, Cayot, P.
   Fast and direct analysis of oxidation levels of oil-in-water emulsions using ATR-FTIR (2019) Food Chem, 293, pp. 307-314.
   34

- Daoud, S, Bou-Maroun, E, Waschatko, G, Horemans, B, Mestdagh, R, Billecke, N
   Detection of lipid oxidation in infant formulas: application of infrared spectroscopy to complex food systems

   (2020) Foods, 9, p. 1432.
   35
- Killeen, DP, Marshall, SN, Burgess, EJ, Gordon, KC, Perry, NB.
   Raman spectroscopy of fish oil capsules: polyunsaturated fatty acid quantitation plus detection of ethyl esters and oxidation (2017) *J Agric Food Chem*, 65, pp. 3551-3558.
   36
- Killeen, DP, Card, A, Gordon, KC, Perry, NB.
   First use of handheld Raman spectroscopy to analyze omega-3 fatty acids in intact fish oil capsules

   (2020) Appl Spectrosc, 74, pp. 365-371.
   37
- Ahmmed, F, Fuller, ID, Killeen, DP, Fraser-Miller, SJ, Gordon, KC.
   Raman and infrared spectroscopic data fusion strategies for rapid, multicomponent quantitation of krill oil compositions

   (2021) ACS Food Sci Technol, 1, pp. 570-578.
   38
- Márquez, C, López, MI, Ruisánchez, I, Callao, MP.
   FT-Raman and NIR spectroscopy data fusion strategy for multivariate qualitative analysis of food fraud

   (2016) *Talanta*, 161, pp. 80-86.
   39
- Borràs, E, Ferré, J, Boqué, R, Mestres, M, Aceña, L, Calvo, A
   Olive oil sensory defects classification with data fusion of instrumental techniques and multivariate analysis (PLS-DA)
   (2016) Food Chem, 203, pp. 314-322.
   40
- Tahir, HE, Xiaobo, Z, Zhihua, L, Jiyong, S, Zhai, X, Wang, S
   Rapid prediction of phenolic compounds and antioxidant activity of Sudanese honey using Raman and Fourier transform infrared (FT-IR) spectroscopy (2017) *Food Chem*, 226, pp. 202-211.
   41
- Fengou, LC, Spyrelli, E, Lianou, A, Tsakanikas, P, Panagou, EZ, Nychas, GJE.
   Estimation of minced pork microbiological spoilage through Fourier transform infrared and visible spectroscopy and multispectral vision technology (2019) *Foods*, 8, p. 238.
   42
- Pacholczyk-Sienicka, B, Ciepielowski, G, Albrecht, Ł.
   The application of NMR spectroscopy and chemometrics in authentication of spices (2021) *Molecules*, 26, p. 382.
   43
- Li, J, Vosegaard, T, Guo, Z.
   Applications of nuclear magnetic resonance in lipid analyses: an emerging powerful tool for lipidomics studies

   (2017) Prog Lipid Res, 68, pp. 37-56.
   44

- Tyl, CE, Brecker, L, Wagner, KH.
   1H NMR spectroscopy as tool to follow changes in the fatty acids of fish oils (2008) *Eur J Lipid Sci Technol*, 110, pp. 141-148.
   45
- Amorim, TL, Granato, ÁS, de Oliveira Mendes, T, de Oliveira, MAL, Amarante, GW, de la Fuente, MA Lipid classification of fish oil omega-3 supplements by 1H NMR and multivariate

**analysis** (2021) *J Food Compos Anal*, 102, p. 104060. 46

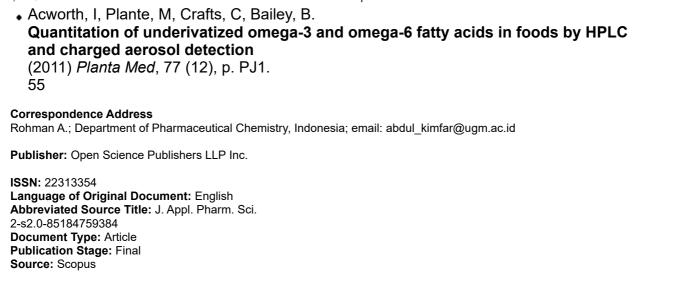
- Lv, J, Wang, C, Zhang, X, Lv, Z, Yu, M.
  1H NMR quantification of DHA and EPA in fish oil (2020) *J Ocean Univ China*, 19, pp. 1193-1197.
  47
- Wu, D, He, Y.
   Potential of spectroscopic techniques and chemometric analysis for rapid measurement of docosahexaenoic acid and eicosapentaenoic acid in algal oil (2014) *Food Chem*, 158, pp. 93-100.
   48
- Wu, D, Chen, X, Cao, F, Sun, DW, He, Y, Jiang, Y.
   Comparison of infrared spectroscopy and nuclear magnetic resonance techniques in tandem with multivariable selection for rapid determination of ω-3 polyunsaturated fatty acids in fish oil (2014) *Food Bioprocess Technol*, 7, pp. 1555-1569.
   49
- Brotas, MSC, Carvalho, GA, Pereira, PAP.
   Determination, through derivatization and GC-MS analysis, of omega-3 and omega-6 fatty acids in fish oil capsules sold in salvador, Bahia
   (2020) *J Braz Chem Soc*, 31, pp. 447-455.
   50
- Abu, EO, Oluwatowoju, I.
   Omega-3 index determined by gas chromatography with electron impact mass spectrometry

   (2009) Prostaglandins Leukot Essent Fat Acids, 80 (4), pp. 189-194.
   51. Apr
- Khedr, A, Alahdal, AM.
   Optimized gas chromatography-mass spectrometric method to profile esterified fatty acids in fish roe and fish oil

   (2018) Indian J Pharm Sci, 80, pp. 628-636.
   52
- Serafim, V, Tiugan, DA, Andreescu, N, Mihailescu, A, Paul, C, Velea, I
   Development and validation of a LC–MS/MS-based assay for quantification of free and total omega 3 and 6 fatty acids from human plasma

   (2019) *Molecules*, 24, p. 360.
   53
- Kotani, A, Watanabe, M, Yamamoto, K, Kusu, F, Hakamata, H.
   Determination of eicosapentaenoic, docosahexaenoic, and arachidonic acids in human plasma by high-performance liquid chromatography with electrochemical detection (2016) Anal Sci, 32, pp. 1011-1014.

```
54
```



# ELSEVIER

Copyright © 2024 Elsevier B.V. All rights reserved. Scopus $^{\mbox{\scriptsize B}}$  is a registered trademark of Elsevier B.V.

*RELX* Group<sup>™</sup>