

Available Online at

http://www.ijcpa.in

April-June 2019

International Journal of CHEMICAL AND PHARMACEUTICAL ANALYSIS

elSSN: 2348-0726 ; plSSN : 2395-2466

DOI: http://dx.doi.org/10.21276/ijcpa

Volume-6

**Review Article** 

Issue-3

Article ID: 0018

## A REVIEW ON EXTRACTION, ANALYSIS AND BIOPOTENTIALOF RUTIN

Harshal Ashok Pawar\*

Assistant Professor, Dr. L. H. Hiranandani College of Pharmacy, Ulhasnagar, Maharashtra, India

\*Corresponding Author: Email: <u>harshal.pawar@dlhhcop.org</u>

Received: 27 May 2018 / Revised: 15 January 2019 / Accepted: 12 June 2019 / Available online: 30 June 2019

### ABSTRACT

Natural compounds such as bioflavonoids have found application in health care system due to their wide biological activities, high safety margins and lower cost. Rutin, a polyphenolic bioflavonoid has shown wide range of therapeutic applications due to its significant antioxidant properties. It has demonstrated several pharmacological activities, including antioxidant, cytoprotective, vasoprotective, anticarcinogenic, neuroprotective and cardioprotective activities. The present review work mainly focused on compilation of the data related to extraction, analysis and biopotential of Rutin from the previously published literatures.

Keywords - Rutin, HPTLC, Isolation, Identification, HPLC Analysis

### 1. INTRODUCTION

Rutin (3,3',4',5,7-pentahydroxyflavone-3-rhamnoglucoside, Fig. 1) is a flavonol, abundantly found in plants, such as passionflower, buckwheat, tea, and apple. It is a vital nutritional component of food stuff<sup>1</sup>. Rutin, also called as rutoside, quercetin-3-rutinoside, and sophorin is a citrus flavonoid glycoside found in buckwheat<sup>2</sup>. The name 'rutin' comes from the plant *Ruta graveolens*, which also contains rutin. Chemically it is a glycoside comprising of flavonolic aglycone quercetin along with disaccharide rutinose. It has demonstrated a number of pharmacological activities, including antioxidant, cytoprotective, vasoprotective, anticarcinogenic, neuroprotective and cardioprotective activities <sup>3-10</sup>.

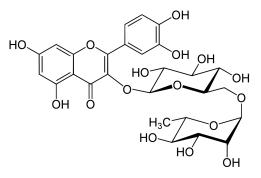


Fig. 1: Chemical structures of Rutin

#### 2. EXTRACTION OF RUTIN

Buckwheat is capable of providing a major dietary source of rutin. Kitabayashi et al. (1995) reported that the rutin content of buckwheat seed ranges from 0.126 to 0.359 mg/g dry weight. Oomah and Mazza (1996) reported 0.47 and 0.77 mg/g dry weight of rutin in whole seed and hulls, respectively. They also reported that flavonoids were highly concentrated in the hulls; the mean flavonoid content of buckwheat seeds and hulls were 3.87 and 13.14 mg/g, respectively.

Prochazka (1985) reported that 6% rutin (wt/wt) was found in carefully dried, *Czechish buckwheat* leaves at the flowering stage. Although most of the details of the industrial production of rutin are proprietary and not described in the open literature, we are aware that Merck GmbH extracts rutin from *fava d'anta* for commercial purposes.

Heywang and Basedow (1992) of Merck GmbH Germany, extracted rutin from shoots of *fava d'anta* (Dimorphandra) with 1,4dioxane under reflux. Rutin was recovered by crystallization at room temperature. Dioxane is, however, considered carcinogenic.

Huo (Chinese Patent 1217329, 1999) described an extraction of rutin from tartary buckwheat seeds by washing with water, coarse grinding, coarse screening, soaking in water, drying in the air, fine grinding, soaking in edible alcohol, extracting below 60°C, and filtering. Balandina et al. (1982) extracted rutin from buckwheat seeds with hot water to remove the desired product and crystallized it.

Zhai (Chinese Patent CN 1160048, 1997) described the extraction of rutin from *Flos sophorae* by soaking with saturated limewater containing 1 - 10% borax and precipitating at pH 1- 6 by adding HC1.

Matsumoto and Hamamoto (1990) recovered rutin from *Sophora augustifolia* buds with methanolic extraction, adsorption onto activated carbon followed by desorption, by elution with 1 % ammonia in 40 % ethanol, and recrystallization from 20 % ethanol.

Liu (1991) described a method of extracting rutin from Japanese Pagoda tree (*Sophora japonicá*) buds by pulverizing, streaming in limewater, neutralizing the supernatant, cooling, filtering, washing, and drying the precipitates. The yield was 14.2% (wt/wt) and the product contained 95.1% (wt/wt) rutin <sup>11</sup>.

### 3. ANALYSIS OF RUTIN

Various analytical methods have been reported in previously published literatures for qualitative and quantitative analysis of Rutin. Various methods have been determined for estimation of rutin and its metabolite such as liquid chromatographic and highperformance liquid chromatography (HPLC) techniques <sup>12</sup>. Recently established method involves infrared spectroscopy, mass spectroscopy, Nuclear magnetic spectroscopy to identify and quantitate rutin <sup>13,14</sup>. Kale and laddha has also been reported for determination of the total flavonoid content and quantitative estimation of rutin by RP-HPLC <sup>15</sup>. Pawar and Salunkhe similarly performed analysis of rutin by U.V. Spectrophotometry <sup>16</sup>. Luftmann et al performed analysis of rutin by mass spectrometry, 1H NMR spectrometry and HPTLC in rat urine. This process requires tedious processing and long time for analysis of sample <sup>17</sup>. Another limitation of existing method includes non-suitability for estimation of rutin in biological samples, as the limit of detection is 50 ng/ml <sup>18</sup>. Apart from above, all the HPLC methods require a long run time for analysis which is based on the retention time of rutin in the mobile phase used. The main problem with rutin is its instability at neutral or basic pH and, also, it is very poor absorption when taken orally or applied topically. Therefore, a precise, reliable, and sensitive method is necessary to estimate rutin in biological fluids <sup>19-22</sup>.

#### 4. **BIOPOTENTIAL OF RUTIN**

Natural compounds such as bioflavonoids have found application in health care system due to their wide biological activities, high safety margins and lower cost. Rutin, a polyphenolic bioflavonoid has shown wide range of therapeutic applications due to its

## International Journal of Chemical & Pharmaceutical Analysis ......April-June 2019

significant antioxidant properties. Rutin possesses several pharmacological activities such as reducing oxidative stress, prevent neuroinflammation, anti-diabetic, aiding to reduce neurodegeneration, cardioprotective activity, wound healing activity, radioprotective activity, nephroprotective activity, hepatoprotective activity, antiplasmodial activity, antiarthritic activity, antiviral activity, improved endothelial functions and antinociceptive activities<sup>23-36</sup>.

### 5. CONCLUSION

Rutin is a common dietary flavonoid that is widely consumed from plant-derived beverages and foods as traditional and folkloric medicine worldwide. Rutin is believed to exhibit significant pharmacological activities, including anti-oxidation, anti-inflammation, anti-diabetic, anti-adipogenic, neuroprotective and hormone therapy. This review has presented a comprehensive overview about the extraction, analysis, and applications of Rutin. The compiled information will be useful to the scientists and researchers for extraction, isolation, identification, and quantitation of Rutin in the plant raw material, crude extracts, and their formulations.

### 6. ACKNOWLEDGEMENT

Author is very much thankful to Dr. Paraag Gide, Principal of Hyderabad Sindhi National Collegiate Boards (HSNCB's) Dr. L. H. Hiranandani College of Pharmacy, Ulhasnagar for his continuous support and encouragement.

### REFERENCES

- 1. Harborne J.B. Nature, distribution, and function of plant flavonoids. Prog. Clin. Biol. Res. 1986; 213:15-24.
- Kreft S., Knapp M., Kreft I. Extraction of rutin from buckwheat (*Fagopyrum esculentum* Moench) seeds and determination by capillary electrophoresis. J. Agric. Food Chem. 1997;47(11):4649–4652.
- Javed H., Khan M.M., Ahmad A., Vaibhav K., Ahmad M.E., Khan A., Ashafaq M., Islam F., Siddiqui M.S., Safhi M.M., Islam F. Rutin prevents cognitive impairments by ameliorating oxidative stress and neuroinflammation in rat model of sporadic dementia of Alzheimer type. Neuroscience. 2012; 17:340–352.
- 4. La Casa C., Villegas I., Alarcón de la Lastra C., Motilva V., Martín Calero M.J. Evidence for protective and antioxidant properties of rutin, a natural flavone, against ethanol induced gastric lesions. J. Ethnopharmacol. 2000;71(1–2):45–53.
- 5. Janbaz K.H., Saeed S.A., Gilani A.H. Protective effect of rutin on paracetamol and CCl<sub>4</sub>-induced hepatotoxicity in rodents. Fitoterapia. 2002; 73:557–563.
- 6. Schwedhelm E., Maas R., Troost R., Böger R.H. Clinical pharmacokinetics of antioxidants and their impact on systemic oxidative stress. Clin. Pharmacokinet. 2003; 42:437–459.
- 7. Trumbeckaite S., Bernatoniene J., Majiene D., Jakstas V., Savickas A., Toleikis A. The effect of flavonoids on rat heart mitochondrial function. Biomed. Pharmacother. 2006; 60:245–248.
- Mellou F., Loutrari H., Stamatis H., Roussos C., Kolisis F.N. Enzymatic esterification of flavonoids with unsaturated fatty acids: effect of the novel esters on vascular endothelial growth factor release from K562 cells. Process Biochem. 2006; 41:2029–3204.
- 9. Nassiri-Asl M., Mortazavi S.R., Samiee-Rad F., Zangivand A.A., Safdari F., Saroukhani S., Abbasi E. The effects of rutin on the development of pentylenetetrazole kindling and memory retrieval in rats. Epilepsy Behav. 2010; 18:50–53.
- 10. Richetti S.K., Blank M., Capiotti K.M., Piato A.L., Bogo M.R., Vianna M.R., Bonan C.D. Quercetin and rutin prevent scopolamine-induced memory impairment in zebrafish. Behav. Brain Res. 2011; 217:10–15.

### International Journal of Chemical & Pharmaceutical Analysis ......April-June 2019

- 11. Chang P, Muir A, inventors; Agri Food Canada AAFC, assignee. Extraction, purification and conversion of flavonoids from plant biomass. United States patent application US 10/528,877. 2006 May 11.
- 12. Morales, M.A., Lozaya, X. Calcium-antagonist effect of rutin on aortic smooth muscle. Planta. Med., 1994; 60: 313-17.
- 13. Sajeeth, C.I., Manna, P.K., Manavalan, R., Jolly, C.I. Quantitative estimation of Gallic Acid, Rutin and Rutin in certain herbal plants by HPTLC method. Der ChemicaSinica., 2010; 1(2): 80-85.
- 14. Boots, A.W., Haenen, G.R., Bast, A. Health effects of rutin: from antioxidant to nutraceutical. Eur J Pharmacol., 2008; 58(5): 325–37.
- 15. Wu, T.H., Yen, F.L., Lin, L.T., Sai, T.R., Lin, C.C., Cham, T.M. Preparation, physicochemical characterization, and antioxidant effects of rutin nanoparticles. Int J Pharm., 2008; 34(6): 160–68.
- 16. Kim, Y.J., Bae, Y.C., Suh, K.T., Jung, J.S. Rutin, a flavonoid, inhibits proliferation and increases osteogenic differentiation in human adipose stromal cells. Bio.Chem.Pharmacol., 2006; 7(2):1268–78.
- 17. Shaji, J., Iyer, S. Preparation, Optimization and In-Vivo Hepatoprotective Evaluation of Rutin Liposomes., Int. J.Curr. Pharm. Res., 2012; 4 (2): 24-32.
- Umathe, S.N., Dixit, P.V., Kumar, V., Bansod, K.U., Wanjari, M.M. Rutin pretreatment increases the bioavailability of pioglitazone in rats: involvement of CYP3A inhibition. BiochemPharmacol., 2008;75:1670–76.
- 19. Chitkara, D., Kumar, S., Mittal, A., Chand, M., Kumar, N. Development of rutin nano formulation and in vivo evaluation using streptozotocin induced diabetic rat model. Drug Del. and Transl. Res., 2012;2:112–23.
- Patil, S.J., Salunkhe, V.R., Alai, M.H. Estimation of Rutin in Ayurvedic Proprietary Medicine by UV-Spectrophotometry., Int. J. Pharm. Sci., 2012; 4(3): 645-47.
- Gupta, N.K., Nahata, A., Dixit, V.K. Development of a spectrofluorimetric method for the determination of curcumin. Asian J.Trad. Med., 2010; 5 (1):12-18.
- 22. Sethiya, N.K., Nahata, A., Dixit, V.K. Simultaneous spectrofluorimetric determination of scopoletin and mangiferin in a methanolic extract of *Canscora decussate* Asian J.Trad. Med., 2008; 3 (6):224-29.
- 23. Javed H, Khan MM, Ahmad A, Vaibhav K, Ahmad ME, Khan A, Ashafaq M, Islam F, Siddiqui MS, Safhi MM. Rutin prevents cognitive impairments by ameliorating oxidative stress and neuroinflammation in rat model of sporadic dementia of Alzheimer type. Neuroscience. 2012 May 17; 210:340-52.
- 24. Hunyadi A, Martins A, Hsieh TJ, Seres A, Zupkó I. Chlorogenic acid and rutin play a major role in the in vivo anti-diabetic activity of Morus alba leaf extract on type II diabetic rats. PloS one. 2012 Nov 21;7(11):e50619.
- 25. Xu PX, Wang SW, Yu XL, Su YJ, Wang T, Zhou WW, Zhang H, Wang YJ, Liu RT. Rutin improves spatial memory in Alzheimer's disease transgenic mice by reducing Aβ oligomer level and attenuating oxidative stress and neuroinflammation. Behavioural brain research. 2014 May 1; 264:173-80.
- 26. Li M, Jiang Y, Jing W, Sun B, Miao C, Ren L. Quercetin provides greater cardioprotective effect than its glycoside derivative rutin on isoproterenol-induced cardiac fibrosis in the rat. Canadian journal of physiology and pharmacology. 2013;91(11):951-9.
- 27. Almeida JS, Benvegnú DM, Boufleur N, Reckziegel P, Barcelos RC, Coradini K, De Carvalho LM, Bürger ME, Beck RC. Hydrogels containing rutin intended for cutaneous administration: efficacy in wound healing in rats. Drug development and industrial pharmacy. 2012 Jul 1;38(7):792-9.
- 28. Patil SL, Mallaiah SH, Patil RK. Antioxidative and radioprotective potential of rutin and quercetin in Swiss albino mice exposed to gamma radiation. Journal of Medical Physics/Association of Medical Physicists of India. 2013 Apr;38(2):87.

# International Journal of Chemical & Pharmaceutical Analysis ......April-June 2019

- 29. Patil SL, Somashekarappa HM, Rajashekhar KP. Radiomodulatory role of Rutin and Quercetin in Swiss Albino mice exposed to the whole-body gamma radiation. Indian journal of nuclear medicine: IJNM: the official journal of the Society of Nuclear Medicine, India. 2012 Oct;27(4):237.
- 30. Kamel KM, Abd El-Raouf OM, Metwally SA, Abd El-Latif HA, El-sayed ME. Hesperidin and rutin, antioxidant citrus flavonoids, attenuate cisplatin-induced nephrotoxicity in rats. Journal of Biochemical and Molecular Toxicology. 2014 Jul;28(7):312-9.
- 31. Khan RA, Khan MR, Sahreen S. CCl 4-induced hepatotoxicity: protective effect of rutin on p53, CYP2E1 and the antioxidative status in rat. BMC complementary and alternative medicine. 2012 Dec;12(1):178.
- 32. Ganesh D, Fuehrer HP, Starzengrüber P, Swoboda P, Khan WA, Reismann JA, Mueller MS, Chiba P, Noedl H. Antiplasmodial activity of flavonol quercetin and its analogues in Plasmodium falciparum: evidence from clinical isolates in Bangladesh and standardized parasite clones. Parasitology research. 2012 Jun 1;110(6):2289-95.
- 33. Horcajada MN, Sanchez C, Scalfo FM, Drion P, Comblain F, Taralla S, Donneau AF, Offord EA, Henrotin Y. Oleuropein or rutin consumption decreases the spontaneous development of osteoarthritis in the Hartley guinea pig. Osteoarthritis and cartilage. 2015 Jan 1;23(1):94-102.
- 34. Carvalho OV, Botelho CV, Ferreira CG, Ferreira HC, Santos MR, Diaz MA, Oliveira TT, Soares-Martins JA, Almeida MR, Junior AS. In vitro inhibition of canine distemper virus by flavonoids and phenolic acids: implications of structural differences for antiviral design. Research in Veterinary Science. 2013 Oct 1;95(2):717-24.
- 35. Ugusman A, Zakaria Z, Chua KH, Megat Mohd Nordin NA, Abdullah Mahdy Z. Role of rutin on nitric oxide synthesis in human umbilical vein endothelial cells. The Scientific World Journal. 2014 Jan 1;2014.
- 36. Selvaraj G, Kaliamurthi S, Thirungnasambandam R, Vivekanandan L, Balasubramanian T. Anti-nociceptive effect in mice of thillai flavonoid rutin. Biomed Environ Sci. 2014 Apr 1;27(4):295-9.