



Analytical Techniques for the Assay of Risperidone: A Review

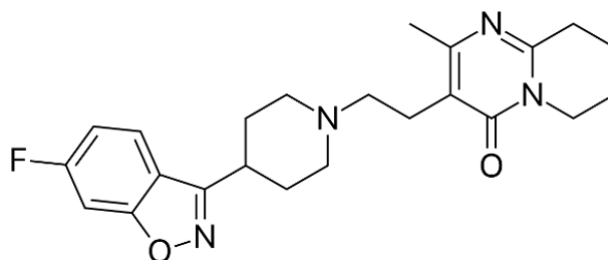
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Annapurna and Sneha Das.****Abstract**

Risperidone is an anti-psychotic drug to reduce this overactivity through inhibition of dopaminergic D2 receptors and serotonergic 5-HT_{2A} receptors in the brain. In the present study the authors have summarised the analytical methods so far published for the estimation of Risperidone in the literature

Keywords: Risperidone; Liquid Chromatographic Method**Introduction**

Risperidone (Figure 1) is a second-generation antipsychotic medication used in the treatment of a number of mood and mental health conditions including schizophrenia and bipolar disorder [1,2]. Chemically it is 3-[2-[4-(6-fluoro-1,2-benzoxazol-3-yl) piperidin-1-yl] ethyl]-2-methyl-6,7,8,9-tetrahydropyrido[1,2-a] pyrimidin-4-one with molecular formula $C_{23}H_{27}FN_4O_2$ and molecular weight 410.5 grams/mole. It is practically insoluble in water (pKa 8.76). It is more soluble in acidic solutions and organic solvents like ethanol and dichloromethane.

Risperidone (CAS no. 106266-06-2) binds with a very high affinity to 5-HT_{2A} receptors, approximately 10-20 fold greater than the drug's binding affinity to D2 receptors, and carries lesser activity at several off-targets which may responsible for some of its undesirable effects. Risperidone is indicated for the treatment of schizophrenia and irritability associated with autistic disorder.

**Figure 1:** Structure of Risperidone ($C_{23}H_{27}FN_4O_2$).

Literature survey reveals that Risperidone was studied by different analytical methods such as spectrophotometry [3-20] (Table 1), Liquid chromatographic methods [22-41] (Table 2) etc for the estimation of Risperidone and. The earlier reported methods were summarized in Table 1.

Table 1: Review of spectrophotometric methods.

Reagent	Linearity ($\mu\text{g/ml}$)	λ_{max} (nm)	Reference
Potassium permanganate in alkaline medium	5-40	415	[3]
Methanol	2-6	280	[4]
Bromo Cresol Green	10-250	440	[5]
MBTH reagent	4-14	666	[6]
0.1N HCl	2.5-20	202	[7]
0.1N HCl: Methanol (40:60)	2-6	280	[8]
0.1N HCl: Methanol (30:70)	3-27	238 & 276	[9]
0.1N HCl	2-6	238	[10]
3% H_2O_2 and Methanol	5-35	274.2	[11]
Britton Robinson Buffer (BR Buffer)	0.11 - 4.93 1.15 -16.82	277 & 238	[12]
Tetrahydrofuran	0.5-5	285	[13]
Water: Acetonitrile (65:35)	21-36	237	[14]
0.1 N HCl	2-12	239	[15]
0.1N NaOH		277	
0.1 N HCl	5-30	238	[16]
Chloramine T(CAT) & Xylene cyanol (XCFF) or Malachite Green (MG)	2-26 2-18	612 619	[17]
7,7,8, 8 Tetra-cyano quinoid methane (TCNQ)	10-110	842	[18]
	8-150	457	
2,3-dichloro-5,6-dicyano-1,4-benzo- quinone (DDQ) tetracyanoethylene (TCNE) Chloranilic acid (CA)	5-70	414	
	25-250	520	
Methanol	10-80	595.8	[19]
Aq. Urea (30%)	4-24	265-287.4	[20]
Phosphate buffer (pH 7.0)	1-50	280	[21]
Acetate buffer (pH 4.0)			
Phosphate buffer (pH 7.5)			

Table 1: Review of liquid chromatographic methods.

Mobile Phase (v/v)	Column	Wavelength (nm)	Flow rate (ml/min)	Linearity (µg/ml)	Reference
Methanol: 0.1M Ammonium Acetate (pH 5.50) (60:40) Chlordiazepoxide hydrochloride (Internal standard)	Supelcosil LC ₈ DB	274	1.0	4.0-275	[22]
Methanol: Acetonitrile (80:20)	Symmetry C18	280	1.0	10-60	[23]
Acetonitrile: 0.2M Potassium Dihydrogen Phosphate (80:20)	Thermo C18	254	0.7	2-10	[24]
0.5% Ammonium Acetate: Acetonitrile: Methanol (50:20:30)	RP-C ₁₈ Phenominex Gemini	260	1.5	20-100	[25]
Methanol: 0.2% aq. OPA (80:20)	Chemisil® ODS C18	235	0.6	0.5-10	[26]
Water: Glacial acetic acid 0.5%: Triethylamine 0.8 %: Acetonitrile (65: 0.32: 0.52: 34.16) Paroxetine (Internal standard)	Purosphere STAR RP-18e	294	-	25-250	[27]
Acetonitrile: 0.05M Potassium dihydrogen phosphate (45:55) (pH 6.5)	C ₁₈	237	1.0	1-100	[28]
Methanol: 0.1% Formic acid (40:60)	Inertsil ODS C-18	260	1.0	4-12	[29]
0.1% Ammonium Formate: Methanol (30:70)	C8	240	1.0	50-3.12	[30]
Methanol: Acetonitrile: 0.02M Phosphate buffer (65:20:15) (pH 3.0)	Hypersil ODS C-18	238	1.0	-	[31]
Methanol: 0.05 M Acetate buffer (pH 4.6): Triethylamine (60:40:0.02)	RP-C18	280	1.0	0.25-10	[32]
Methanol: Acetonitrile: 50 mM Potassium Dihydrogen Ortho phosphate (80:10:10)	Phenomenex Gemini C-18	234	1.3	0.5-0.99	[33]
Water: Acetonitrile (75:25)	Hypersil BODS C-18	280	0.8	40-80	[34]
10 mM potassium dihydrogen phosphate, pH 3.5 ± 0.05): acetonitrile: methanol (65:20:15)	Waters Xterra RP8	276	1.0	4.973-44.757	[35]
Methanol: Acetonitrile: Potassium dihydrogen ortho-phosphate (60:30:10)	C18	234	1.0	20-100	[36]
Water: Methanol (35:65) (pH 5.5)	Phenomenex C18	276	0.9	10-100	[37]
Acetonitrile: Methanol (20:80)	Capcell Pak C18	239	1.2	20-100	[38]
Ammonium acetate buffer (pH 6.8): Acetonitrile	RP-18	260	0.5	0.5-3	[39]
Methanol: Acetonitrile (75:25)	Chromosil C18	238	1.0	20-100	[40]
Methanol: Ammonium Formate (pH 3.0 with OPA) (90:10)	C18	254	1.0	20-100	[41]

Conclusion

This review article explains different analytical methods developed for the estimation of Risperidone in pharmaceutical dosage forms as well as biological fluids.

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