

**ANALYTICAL TECHNIQUES FOR THE ASSAY OF ETORICOXIB - A
REVIEW**

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ABSTRACT

Etoricoxib is an oxicam-class nonsteroidal anti-inflammatory drug. An enzyme called cyclooxygenase-2 (COX-2) is responsible for pain and inflammation. Etoricoxib is the most recent selective (COX-2) inhibitor with more COX-2 selectivity than previous COX-2-selective NSAIDs, significantly improving its gastric safety profile. 5-Chloro-6'-methyl-3-[4-(methylsulfonyl)phenyl]-2,3'-bipyridine is the IUPAC name of Etoricoxib. Etoricoxib is water insoluble but rapidly soluble in alkaline aqueous solutions. The current therapeutic indications of etoricoxib include treating several painful conditions, such as osteoarthritis, acute gout, ankylosing spondylitis, and rheumatoid arthritis. The following review covers the analytical techniques for the analysis of Etoricoxib.

KEYWORD: Etoricoxib, Method development, Cyclooxygenase-2, Nonsteroidal anti-inflammatory drugs, pain.

INTRODUCTION

Selective cyclooxygenase (COX)-2 inhibitors are a class of nonsteroidal anti-inflammatory drugs (NSAIDs) that have become important in the pharmacological management of pain and arthritis.^[1] Etoricoxib (ETX) chemically 5-chloro-2-(6-methylpyridin-3-yl)-3-(4-methylsulfonylphenyl) pyridine is a methyl sulphone derivative (Figure 1).^[2,3] Etoricoxib is also a bipyridine derivative. This newer COX-2 inhibitor has the highest COX-2 selectivity.^[4]

Etoricoxib is generally administered through an oral route. Etoricoxib (Figure 1) has a molecular formula $C_{18}H_{15}ClN_2O_2S$ and a molecular weight 358.8419 g/mole, and it is a white or off-white powder with a bitter taste. The pKa values of Etoricoxib are found to be 3.9 and 4.6.^[5,6]

Etoricoxib selectively inhibits isoform 2 of the enzyme cyclooxygenase (COX-2), it is mainly used in the management of^[1]

- Osteoarthritis,
- Rheumatoid Arthritis,
- Acute Gouty Arthritis,
- Ankylosing Spondylitis
- Low Back Pain,
- Acute Postoperative Pain,
- Primary Dysmenorrhea

Side effects are^[4]

- Dyspepsia,
- Abdominal Pain,
- Pedal Edema,
- Rise in Blood pressure
- Dry Mouth,
- Aphthous Ulcers,
- Taste Disturbance And
- Paresthesias

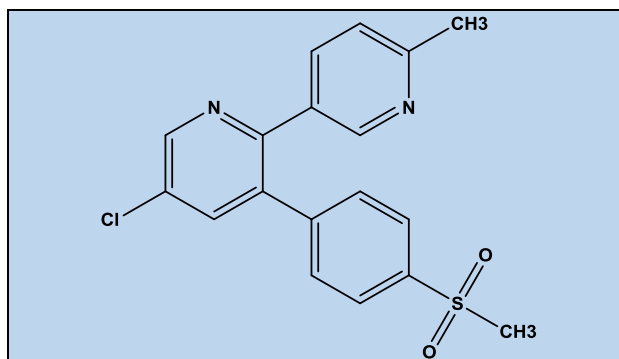


Figure 1: Chemical structure of Etoricoxib.

The present review article summarizes the analytical techniques so far developed, such as spectrophotometry^[7-14] (Table 1), high performance thin layer chromatography^[15-16] (Table 2), high-performance liquid chromatography^[17-28] (Table 3), and Hyphenated techniques^[29-30] (Table 4) for the determination of etoricoxib and some of the analytical parameters were highlighted.

1. Spectrophotometric method

Table 1: Determination of Etoricoxib (ETX) by UV Spectrophotometric Method.

Solvent/Reagent	λ_{max} (nm)	Linearity ($\mu\text{g/ml}$)	Reference
0.1N HCl	233	2-24	[7]
0.1N HCl	234	3-15	[8]
Methanol and phosphate buffer (pH 7.2)	284	4-24	[9]
0.1 N HCl	233	0.1-0.5	[10]
Methanol	248	1-8	[11]
Chloroform	247	1-40	[12]
Methanol	234	1-11	[13]
Methanol	284	5-50	[14]

2. High Performance Thin Layer Chromatography (HPTLC)

Table 2: Determination of etoricoxib (ETX) by HPTLC Method.

Mobile Phase	λ (nm)	Stationary Phase	Linearity Range (ng/band)	LOD (ng/band)	LOQ (ng/band)	Reference
toluene: ethyl acetate: methanol (6: 4: 1)	258	silica gel 60 F	50-300	-	-	[15]
ethyl acetate-methanol (8:2)	290	silica gel 60F254	50-250	10.993	33.314	[16]

3. High Performance Liquid Chromatography (HPLC)

Table 3: Determination of etoricoxib (ETX) by HPLC Method.

Technique	Mobile Phase	Flow rate	LOD ($\mu\text{g/ml}$)	LOQ ($\mu\text{g/ml}$)	References
RP-HPLC	Acetonitrile: pH 3 phosphate buffer (70:30% v/v)	1.0 ml/min	3.3172	8.132	[17]
RP-HPLC	Acetonitrile: Water (55:45 %v/v)	0.8 ml/min	0.0837	0.253	[18]
RP-HPLC	mixture of solution (1 ml TFA in 2-liter milli-Q water) and acetonitrile (75:25 v/v).	1.5 ml/min	0.5273	1.598	[19]
RP-HPLC	mixture of phosphate buffer (PH6, adjusted with orthophosphoric acid) and methanol (30:70 v/v)	1.2 ml/min	0.0115	0.0348	[20]

RP-HPLC	acetonitrile and potassium dihydrogen phosphate buffer (pH 4.2) (46:54 % v/v)	1.2 ml/min	0.0704	0.2134	[21]
RP-HPLC	acetonitrile: (0.05M) KH ₂ PO ₄ buffer (50:50)	1.8 ml/min	0.193	0.450	[22]
HPLC	mixture of acetonitrile: methanol: 10mM potassium dihydrogen phosphate (pH 3.0 adjusted with orthophosphoric acid)	1 ml/min	-	-	[23]
HPLC	ammonium acetate buffer: acetonitrile (65:35 v/v)	1 ml/min	10	20	[24]
HPLC	Methanol	1 ml/min	250	650	[25]
RP-UFLC	Acetonitrile: water (50: 50)	0.8 ml/min	0.0291	0.0894	[26]
RP-HPLC	methanol, and phosphate buffer of pH 6 in a ratio of 70:30 (v/v)	0.8 ml/min	5.08	16.94	[27]
RP-HPLC	acetonitrile, methanol and water in the proportion of 60:15:25 (v/v/v)	1 ml/min	-	-	[28]

4. Hyphenation Techniques

Table 4: Determination of etoricoxib (ETX) by hyphenated Method.

Mobile Phase	Column	Method	Flow Rate	LOQ (ng/ml)	Reference
component A, 5mM KH ₂ PO ₄ adjusted to pH 2.5 with 5mM H ₃ PO ₄ , and component B, acetonitrile.	LC-MS AQ-ODS	isocratic elution	1 ml/min	0.04	[29]
0.01M acetate buffer pH 5.0 -acetonitrile (60 : 40, v/v)	UPLC TM BEH C18	gradient elution	0.3 ml/min	-	[30]

CONCLUSION

Various analytical techniques for estimating Etoricoxib in bulk, pharmaceutical formulations, and biological samples were reported, including UV, HPTLC, HPLC, UPLC and LC-MS methods. This review will help readers in understanding the analytical methodologies presented for quantifying Etoricoxib.

CONFLICT OF INTEREST: There isn't any potential for a conflict of interest.

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