

|         |  |    |
|---------|--|----|
| 1.7.1.3 | The Transporters of Intestinal   | 39 |
| 1.8     | Pre-Clinical Studies   | 40 |
| 1.9.    | Method Validation  | 40 |
| 1.9.1   | Precision  | 41 |
| 1.9.2   | Accuracy   | 41 |
| 1.9.3   | Linearity  | 42 |
| 1.9.4   | Range  | 42 |
| 1.9.5   | Ruggedness   | 42 |
| 1.9.6   | Limit of Detection   | 43 |
| 1.9.7   | Limit of Quantitation  | 43 |
| 1.9.8   | Selectivity  | 43 |
| 1.9.9   | Specificity  | 43 |
| 1.9.10  | Stability  | 43 |
| 1.10.   | Internal Standard  | 44 |
| 1.11.   | Previous Analytical Studies and Literature Survey                      | 46 |
| 1.12.   | Objective of This Study  | 55 |
|         | <b>Chapter Two: Experimental Part</b>                                  | 56 |
| 2       | Experimental Part  | 57 |
| 2.1     | Reagents   | 57 |
| 2.2     | Instrumentation  | 58 |
| 2.3     | Animals  | 58 |
| 2.4     | Preparation of Stock Solutions   | 59 |
| 2.4.1   | Preparation of Candesartan Solution to be Given to the Rats            | 59 |
| 2.4.2   | Preparation of Stock Solution of Candesartan                           | 60 |
| 2.4.3   | Preparation of Stock Solution for Internal Standard                    | 60 |
| 2.4.4   | Preparation of 2 ng/ml Irbesartan IS in methanol (precipitating agent) | 60 |
| 2.4.5   | Preparation of Working Solution for Candesartan                        | 60 |
| 2.4.5   | Preparation of Candesartan STD Serial Dilution and Spiked Plasma       | 60 |
| 2.4.6   | Preparation of Candesartan QC Serial Dilution and Spiked Plasma        | 60 |
| 2.5     | Preparation of Orange, Licorice and Pomegranate Juices                 | 62 |
| 2.6     | Method of Sample Preparation   | 62 |
| 2.7     | Validation   | 63 |
| 2.7.1   | Accuracy and Precision   | 63 |
| 2.7.2   | Specificity and Selectivity  | 63 |
| 2.7.3   | Sensitivity  | 63 |