

# Index

Agradecimientos.....	5
List of Figures .....	13
Acronyms .....	21
Abstract .....	23
Resumen .....	25
Resum.....	27
Chapter 1 .....	29
1.1. Introduction and motivation.....	29
1.2. Objectives.....	31
1.3. Structure of the thesis.....	32
Chapter 2 .....	35
State of the art.....	35
2.1. Introduction to the heart.....	35
2.1.1. Cardiac Action Potential.....	37
2.1.2. Cardiac Electrophysiology .....	38
2.1.3. Atrial Fibrillation.....	38
2.2. The electrocardiogram.....	46
2.3. Body Surface Potential Mapping .....	47
2.3.1. Body Surface Potential Mapping in atrial fibrillation .....	48
2.4. Electrocardiographic Imaging .....	49
2.4.1. Forward Problem of electrocardiography.....	50
2.4.2. Inverse Problem of electrocardiography .....	51
2.4.3. ECGI for atrial fibrillation.....	53
2.5. Cardiac Modelling.....	56
2.5.1. Cellular level modelling .....	56
2.5.2. Atrial cell mathematical models.....	57
2.5.3. Tissue level modelling.....	58
2.5.4. Geometrical models.....	60
Compendium of articles .....	63

Chapter 3 .....	65
Effects of Torso Mesh Density and Electrode Distribution on the Accuracy of Electrocardiographic Imaging During Atrial Fibrillation.....	65
Abstract .....	65
3.1. Introduction .....	66
3.2. Materials and methods .....	67
3.2.1. Study population - Data acquisition .....	67
3.2.2. Data processing .....	68
3.2.3. Quality of mesh evaluation metrics.....	71
3.2.4. Frequency metrics .....	71
3.3. Results.....	71
3.3.1. Impact of mesh density on ECGI reconstruction .....	71
3.3.2. Impact of electrode relocation in low-density torso meshes on ECGI reconstruction.....	74
3.4. Discussion .....	77
3.4.1. Limitations .....	79
3.5. Conclusion.....	80
3.6. Acknowledgements .....	80
References.....	81
Chapter 4 .....	85
Robustness of Imageless Electrocardiographic Imaging Against Uncertainty in Atrial Morphology and Location .....	85
Abstract .....	85
4.1. Introduction .....	86
4.2. Material and Methods.....	87
4.3. Results .....	89
4.4. Discussion .....	91
4.5. Conclusion.....	92
4.6. Acknowledgments.....	92
References.....	92

Chapter 5 .....	95
Improving Electrocardiographic Imaging Solutions: A Comprehensive Study on Regularization Parameter Selection in L-curve Optimization.....	95
Abstract .....	95
5.1. Introduction .....	96
5.2. Material and methods .....	97
5.2.1. Cardiac simulations .....	97
5.2.2. Patient data .....	98
5.2.3. Modelling the influence of noise.....	99
5.2.4. Inverse problem calculation .....	100
5.3. Results .....	102
5.3.1. Effect of noise in the inverse problem using atrial simulations .....	102
5.3.2. Patient examples.....	108
5.4. Discussion .....	112
5.4.1. Effects of Noise in the L-curve .....	112
5.4.2. Effects of noise in the extracted ECGI maps.....	113
5.4.3. Limitations .....	114
5.5. Conclusion.....	114
Acknowledgements .....	114
References.....	115
Chapter 6 .....	119
Filtering Strategies of Electrocardiographic Imaging Signals for Stratification of Atrial Fibrillation Patients .....	119
Abstract .....	119
6.1. Introduction .....	120
6.2. Methods.....	121
6.2.1. Patient signal and geometry acquisition .....	121
6.2.2. ECGI post-processing .....	122
6.2.3. Reentrant activity detection.....	122
6.2.4. Reentrant activity evaluation and statistical analysis .....	123
6.3. Results .....	124
6.3.1. Reentrant activity analysis.....	124
6.3.2. Post-processing effects and PVI outcome .....	130

6.4.	Discussion .....	131
6.4.1.	ECGI-derived phase metrics and PVI outcome.....	132
6.4.2.	Effects of filtering ECGI signals for rotor detection .....	133
6.4.3.	Effects of time-space criteria for rotor detection.....	133
6.4.4.	Clinical implications .....	134
6.4.5.	Limitations and future work .....	135
6.5.	Conclusion.....	135
	Funding Sources.....	136
	References.....	136
	Chapter 7 .....	139
	Higher Reproducibility of Phase Derived Metrics from Electrocardiographic Imaging During Atrial Fibrillation in Patients Remaining in Sinus Rhythm after Pulmonary Vein Isolation .....	139
	Abstract .....	139
7.1.	Introduction .....	140
7.2.	Methods.....	141
7.2.1.	Study Population .....	141
7.2.2.	Data acquisition.....	142
7.2.3.	Data processing .....	142
7.2.4.	Atrial fibrillation complexity quantification .....	143
7.2.5.	Reproducibility measurements.....	143
7.2.6.	Statistical analysis .....	143
7.2.7.	Outcome prediction based on ECGI reproducibility .....	144
7.3.	Results .....	144
7.3.1.	Reproducibility of ECGI metrics vs. patient outcome .....	145
7.3.2.	ECGI Reproducibility vs. AF type .....	148
7.3.3.	Association of PVI success based on ECGI variability metrics.....	149
7.4.	Discussion .....	150
7.4.1.	Mechanism of AF and PVI outcome .....	150
7.4.2.	Temporal reproducibility of ECGI derived metrics.....	151
7.4.3.	Clinical implications .....	151
7.5.	Limitations .....	151
7.6.	Conclusions .....	152
	Acknowledgments.....	152

Funding Sources.....	152
References.....	153
Chapter 8 .....	157
Complexity and Recurrence of Body Surface Electrocardiograms Correlates with Estimated Reentrant Atrial Activity with Electrocardiographic Imaging in Atrial Fibrillation Patients .....	157
Abstract .....	157
8.1. Introduction.....	158
8.2. Methods.....	159
8.2.1. Data acquisition and processing.....	159
8.2.2. Assessment of AF substrate complexity and AF propagation recurrence on Body Surface Potential Mapping.....	160
8.2.3. Reentrant atrial activity analysis from ECGI signals .....	162
8.2.4. Statistical analysis .....	162
8.3. Results.....	163
8.3.1. BSPM-based versus ECGI-based AF substrate complexity and AF propagation recurrence .....	163
8.3.2. BSPM-based AF substrate complexity and AF propagation recurrence vs ECGI-based reentrant atrial activity .....	164
8.3.3. Assessment of left and right atrium differences.....	165
8.3.4. BSPM-based AF substrate complexity and AF propagation recurrence vs ECGI-based reentrant atrial Activity per atrium.....	167
8.4. Discussion .....	169
8.4.1. Relationship between reentrant activity and BSPM-based AF complexity and AF propagation recurrence.....	170
8.4.2. Left and right atrium assessment of AF substrate complexity.....	170
8.4.3. Limitations .....	171
8.5. Conclusion.....	171
Acknowledgments.....	172
Supplementary Table.....	172
References.....	173
Chapter 9 .....	177
Discussion and conclusions .....	177
9.1. Main findings .....	177
9.2. Comparison with previous studies .....	179

9.3.	Limitations .....	180
9.4.	Conclusions .....	181
9.5.	Guidelines for future works.....	184
Chapter 10 .....		185
Contributions .....		185
10.1.	Main contributions of this thesis .....	185
10.1.1.	Journal Papers .....	185
10.1.2.	International Conferences.....	186
10.2.	Contributions related to this thesis .....	187
10.2.1.	Journal Papers .....	187
10.2.2.	International Conferences.....	187
10.2.3.	National Conferences .....	188
10.3.	Patents .....	188
10.4.	Awards.....	188
10.5.	Participation in scientific international committees .....	188
10.6.	Diffusion of results.....	189
10.7.	Industrial collaboration .....	189
10.8.	Teaching .....	189
10.8.1.	Supervision of bachelor thesis.....	189
10.8.2.	Supervision of master thesis.....	190
10.9.	Research stay .....	190
10.10.	Research projects and funding .....	190
References.....		193