

## S2 TEXT: Q10 VALUES

### **Modeling hypothermia induced effects for the heterogeneous ventricular tissue from cellular level to the impact on the ECG**

<sup>1</sup>Roland Kienast, <sup>1</sup>Michael Handler, <sup>1</sup>Markus Stöger, <sup>1,3</sup>Daniel Baumgarten, <sup>1</sup>Friedrich Hanser, <sup>1,2</sup>Christian Baumgartner

<sup>1</sup> Institute of Electrical and Biomedical Engineering, UMIT – University for Health Sciences, Medical Informatics and Technology, A-6060 Hall in Tyrol, Austria

<sup>2</sup> Institute of Health Care Engineering with European Testing Center of Medical Devices, Graz University of Technology, A-8010 Graz, Austria

<sup>3</sup>Institute of Biomedical Engineering and Informatics, Technische Universität Ilmenau, D-98693 Ilmenau, Germany

### **Content**

Q10 values extracted from literature.....	2
APD .....	2
V <sub>max</sub> .....	2
CL .....	3
CV .....	3
References.....	4

## Q10 values extracted from literature

This supplemental includes estimated  $Q_{10}$  values based on literature where the  $Q_{10}$  values (APD,  $V_{\max}$ , CL, CV) are calculated based on two values for different temperatures, respectively, using

$$Q_{10} = \left( \frac{X_{\text{high}}}{X_{\text{low}}} \right)^{\left( \frac{10}{T_{\text{high}} - T_{\text{low}}} \right)} \quad (5)$$

where  $X_{\text{high}}$  denotes the value of the investigated feature at higher temperature  $T_{\text{high}}$  and  $X_{\text{low}}$  the value at the lower temperature  $T_{\text{low}}$ . Thereby, the two values at high and low temperature are extracted from textual description or estimated based on available graphics in the respective literature. Table I summarizes the results for APD, Table II for  $V_{\max}$ , Table III for CL and Table IV for CV.

### APD

Table I: Estimated  $Q_{10}$  values for APD based on literature

$Q_{10}$	Measure	Species	Tissue	Temperature range	Reference
0.66	APD90	guinea-pig	ventricular papillary muscle	37-27	[1]
0.83	APD95	rabbit	left ventricle	33-23	[2]
0.53	APD90	rabbit	isolated papillary muscles	37-27	[3]
0.44	APD90	guinea-pig	ventricular papillary muscle	37-27	[4]
0.48	APD90	guinea-pig	ventricular and atrial cells	35-25	[5]
0.75	APD90	pig	ventricular septum	37-32	[6]
0.61	APD70	rabbit	ventricle	37-17	[7]
0.58	APD	dog	left ventricle	36-32	[8]
0.55	APD90	rats	papillary muscles	37-10	[9]

### $V_{\max}$

Table II: Estimated  $Q_{10}$  values for  $V_{\max}$  based on literature

$Q_{10}$	Species	Tissue	Temperature range	Reference
1.56	guinea-pig	ventricular papillary muscle	37-27	[1]
1.51	chicken	right ventricle	37-20	[10]
1.59	rabbit	left ventricle	33-23	[2]
1.65	rabbit	isolated papillary muscles	37-27	[3]
1.4	guinea-pig	ventricular papillary muscle	37-27	[4]
2.35	guinea-pig	ventricular cells	35-25	[5]

The obtained  $Q_{10}$  value for  $V_{\max}$  of 1.68 used in the model was estimated by calculating the mean value of all listed  $Q_{10}$  values in table II.

## CL

In literature, the information about heart rate is usually indicated by beats per minute (ppm). To convert values from ppm to CL following formula was used

$$CL = \frac{60}{ppm}$$

where CL denotes the value for the cycle length in seconds and ppm the value for the heart rate in beats per minute.

Table III: Estimated  $Q_{10}$  values for CL based on literature

$Q_{10ppm}$	$Q_{10CL}$	Species	Temperature range	Reference
3	0.35	rat	34-23	[11]
2	0.5	pig	37-31	[12]
3.6	0.28	dog	40-20.5	[13]

## CV

Table IV: Estimated  $Q_{10}$  values for CV based on literature

$Q_{10}$	Species	Tissue	Temperature range	Reference
1.5	chicken	right ventricle	37-20	[10]
1.39	rabbit	ventricle	37-27	[7]
1.47	rabbit	ventricle	37-27	[14]
1.59	dog	left ventricle	36-26	[15]

## References

1. Melnikov AL, Lathrop DA, Helgesen KG. Diazepam-induced Ca(2+)-channel blockade reduces hypothermia-induced electromechanical changes in isolated guinea pig ventricular muscle. *Eur J Anaesthesiol.* 1998;15(1):96–102. PubMed PMID: 9522148. eng.
2. Samson JP, Reisin I, Ruiz-Ceretti E, Schanne OF. Effects of low temperature on intracellular ionic concentrations and transmembrane potential in isolated rabbit hearts. *J Mol Cell Cardiol.* 1977;9(1):39–50. PubMed PMID: 845974. eng.
3. Fedorov VV, Li L, Glukhov A, Shishkina I, Aliev RR, Mikheeva T, et al. Hibernator *Citellus undulatus* maintains safe cardiac conduction and is protected against tachyarrhythmias during extreme hypothermia: possible role of Cx43 and Cx45 up-regulation. *Heart Rhythm.* 2005;2(9):966–75. doi: 10.1016/j.hrthm.2005.06.012. PubMed PMID: 16171752. eng.
4. Bjørnstad H, Tande PM, Lathrop DA, Refsum H. Effects of temperature on cycle length dependent changes and restitution of action potential duration in guinea pig ventricular muscle. *Cardiovasc Res.* 1993;27(6):946–50. PubMed PMID: 8221783. eng.
5. Hume JR, Uehara A. Ionic basis of the different action potential configurations of single guinea-pig atrial and ventricular myocytes. *J Physiol.* 1985;368:525–44. PubMed PMID: 2416918. eng.
6. Roscher R, Arlock P, Sjöberg T, Steen S. Effects of dopamine on porcine myocardial action potentials and contractions at 37 degrees C and 32 degrees C. *Acta Anaesthesiol Scand.* 2001;45(4):421–6. PubMed PMID: 11300379. eng.
7. Egorov YV, Glukhov AV, Efimov IR, Rosenshtraukh LV. Hypothermia-induced spatially discordant action potential duration alternans and arrhythmogenesis in nonhibernating versus hibernating mammals. *Am J Physiol Heart Circ Physiol.* 2012;303(8):H1035-46. doi: 10.1152/ajpheart.00786.2011. PubMed PMID: 22886418. eng.
8. Piktel JS, Rosenbaum DS, Wilson LD. Mild hypothermia decreases arrhythmia susceptibility in a canine model of global myocardial ischemia\*. *Crit Care Med.* 2012;40(11):2954–9. doi: 10.1097/CCM.0b013e31825fd39d. PubMed PMID: 22890250. eng.
9. Wang S-Q, Cao H-M, Zhou Z-Q. Temperature dependence of the myocardial excitability of ground squirrel and rat. *J Therm Biol.* 1997;22(3):195–9. doi: 10.1016/S0306-4565(97)00010-7.
10. Hirota A, Fujii S, Sakai T, Kamino K. Temperature dependence of spontaneous electrical activity in early embryonic heart monitored optically with a potential-sensitive dye. *Jpn J Physiol.* 1983;33(1):85–100. PubMed PMID: 6855032. eng.
11. Harary I, Farley B. In vitro studies on single beating rat heart cells. II. Intercellular communication. *Exp Cell Res.* 1963;29:466–74. PubMed PMID: 13952709. eng.
12. Weisser J, Martin J, Bisping E, Maier LS, Beyersdorf F, Hasenfuss G, et al. Influence of mild hypothermia on myocardial contractility and circulatory function. *Basic Res Cardiol.* 2001;96(2):198–205. PubMed PMID: 11327339. eng.
13. Berne RM. Myocardial Function in Severe Hypothermia. *Circ Res.* 1954;2(1):90–5. doi: 10.1161/01.RES.2.1.90.
14. Fedorov VV, Glukhov AV, Sudharshan S, Egorov Y, Rosenshtraukh LV, Efimov IR. Electrophysiological mechanisms of antiarrhythmic protection during hypothermia in winter hibernating versus nonhibernating mammals. *Heart Rhythm.* 2008;5(11):1587–96. doi: 10.1016/j.hrthm.2008.08.030. PubMed PMID: 18984537. eng.
15. Piktel JS, Jeyaraj D, Said TH, Rosenbaum DS, Wilson LD. Enhanced dispersion of repolarization explains increased arrhythmogenesis in severe versus therapeutic hypothermia. *Circ Arrhythm Electrophysiol.* 2011;4(1):79–86. doi: 10.1161/CIRCEP.110.958355. PubMed PMID: 21163888. eng.