## Effects of morphological and functional heterogeneities on the intracellular $Ca^{2+}$ signals in coupled pancreatic $\beta$ -cells

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The Islets of Langerhans are mainly composed of insulin-secreting pancreatic  $\beta$ -cells, glucagon-secreting  $\alpha$ -cells and somatostatin-secreting  $\delta$ -cells[1]. At the cellular level, secretion of these hormones takes place through a common mechanism involving glucose metabolism, electrical activity and Ca<sup>2+</sup>-handling[2]. In addition, pancreatic hormone secretion is regulated by intra-islet interactions including paracrine and autocrine signals, as well as electrical coupling mediated by gap junctions between  $\beta$ -cells[3].

Electrical coupling between  $\beta$ -cells has been previously studied both theoretically and experimentally. In these studies, it was shown that  $\beta$ -cell coupling is essential for the synchronized release of insulin[1]. In addition, it was demonstrated that the lack of functional gap junctions leads to impaired pulsatile insulin secretion due to uncoordinated Ca<sup>2+</sup> oscillations[4].

In this work we used a computational model to assess the effect of morphological and functional heterogeneities in the islet  $\beta$ -cells (including differences in cell sizes,  $\beta$ -cell interconnectivity and electrophysiological and Ca<sup>2+</sup> buffering properties) on the Ca<sup>2+</sup> signal produced in the cytosol, ultimately related to the secretory response of the islet  $\beta$ -cells.

## References

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