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Bio-based ionic liquids as extraction-preservation strategies in RNA bioprocessing

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PURPOSE OF THE ABSTRACT

Nucleic acids are powerful biological tools in several fields [1], ranging from fundamental to applied research. However, the ubiquitous instability of ribonucleic acid (RNA) along with issues associated to its purity degree have been preventing its widespread use as low-cost biotherapeutics. Based on the well-known capacity of amino acids to specifically interact with RNA when used as ligands in chromatography [2], a set of amino-acid-based ionic liquids (AA-ILs) was herein investigated, both to act as preservation media and as phase-forming agents of aqueous biphasic systems (ABS) to carry out the biopolymer extraction from real matrices. AA-ILs comprising the cholinium cation and anions derived from L-lysine ([Ch][Lys]), L-arginine ([Ch][Arg]), L-glutamic acid ([Ch][Glu]) and DL-aspartic acid ([Ch][Asp]) were studied for this purpose. This work proved that the stability of RNA is preserved in aqueous solutions of AA-ILs, even in presence of ribonucleases (RNases). It is also demonstrated that these AA-ILs display no cytotoxicity onto two distinct human cell lines. After ensuring the stability of RNA in presence of AA-ILs and protection against RNases, ABS formed by AA-ILs were applied in the extraction of RNA from a bacterial lysate sample, showing that RNA can be successfully extracted to the IL-rich phase, which can also act as a preservation medium. RNA can be recovered by alcohol precipitation and the IL reused, contributing to the development of an integrated and sustainable extraction-preservation process for nucleic acids toward the current critical demand of high-quality/high-purity biotherapeutics.

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FIGURE 1

FIGURE 2

KEYWORDS

ribonucleic acid | ionic liquids | aqueous biphasic systems | integrated processes

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