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Title: Use of designed experiments for the improvement of pre-analytical workflow for the quantification of intracellular nucleotides in cultured cell lines

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2 **Highlights:**

- 3 • Pre-analytical workflow is a critical step for intracellular analysis
- 4 • Approach with designed experiments was used
- 5 • The optimal organic solvent used for lysis and extraction is cell line dependent
- 6 • Methanol/water (70/30, v/v) is a good compromise for nucleotide extraction

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