Supplementary information

Supplementary information S1. IUPAC recommendations to calculate the sensor performance parameters:

1. Linear range:

The linear range of the electrode is defined as that part of the calibration curve through which a linear regression demonstrates no deviation from linearity by more than 2 mV.

2. Limit of detection (LOD):

The concentration of the primary ion at the point of intersection of the extrapolated lines of the Nernstian (high concentration) and nonresponsive (low concentration) segments of the calibration curve can be considered as an "attainable" detection limit under the stated experimental conditions.

3. Dynamic response time:

Practical response time required for the selected sensors to reach a value within ± 1 mV of the final stabilized potential upon increasing the concentration of EPH.

4. Reversibility:

The equilibrium potential was alternatively measured in 1.56×10^{-4} and 1.57×10^{-3} M ephedrine solutions.

Supplementary information S2. Derivation of the standard addition equation (equation 2).

Equation 2 was derived from Nernst equation as following:

- The measured potential in the sample solution of unknown concentration C_u and known volume V_s .

$$E_1 = K + s \log C_s$$

Equation 1

Where (K) is the intercept, (s) is the slope and (C_s) is the sample molar concentration.

- The measured potential in the sample solution after addition of known volume V_{st} of standard EPH solution of molar concentration C_{st} .

$$E_2 = K + s\log \frac{C_s V_s + C_{st} V_{st}}{V_t}$$

Equation 2

- V_t is the total volume of the sample (V_s) and the added standard (V_{st}) solutions and S is the slope measured in the blank matrix.

$$\Delta E = E_2 - E_1 = s \log \frac{C_s V_s + C_{st} V_{st}}{C_s V_t}$$
Equation 3
$$10^{\left(\frac{\Delta E}{S}\right)} = \frac{C_u V_s + C_{st} V_{st}}{C_s V_t}$$
Equation 4

Equation 4

$$[C_{s}V_{t} \times 10^{\left(\frac{\Delta E}{S}\right)}] - C_{s}V_{s} = C_{st}V_{st}$$
 Equation 5

$$C_{s}\left[V_{t} \times 10^{(-/S)} - V_{s}\right] = C_{st}V_{st}$$
 Equation 6

$$C_{s}(M) = \frac{C_{st}V_{st}}{\left[V_{t} \times 10^{\left(\frac{\Delta E}{S}\right)} - V_{s}\right]}$$
Equation 7

 $C_{s}(g/L) = \frac{C_{st}V_{st}}{\left[V_{t} \times 10^{\left(\Delta E/S\right)} - V_{s}\right]} \times Mwt$

Equation 8

$$C_{s}(g/g) = \frac{C_{st}V_{st}}{\left[V_{t} \times 10^{\left(\frac{\Delta E}{S}\right)} - V_{s}\right]} \times \frac{Mwt}{w} \times \frac{200}{1000}$$

Equation 9

$$C_{s}(ppm) = \frac{C_{st}V_{st}}{\left[V_{t} \times 10^{\left(\frac{\Delta E}{S}\right)} - V_{s}\right]} \times \frac{Mwt \times 200 \times 10^{6}}{1000 \times w}$$

Equation 10

$$\frac{3.3048 \times 10^7 \times C_{st} V_{st}}{W \times \left[\left(V_t \times 10^{\left(\frac{\Delta E}{s} \right)} \right) - V_s \right]}$$

For Ephedrine: Equation 11

Supplementary information S3. Post hoc statistical analysis within levels of statistically significant factors.

Dognongo	Ion Exchanger (<i>P</i> value)					
Response		AR	PM	РТ	TPB	
	AR					
Slope	PM	<u>0.0312</u>				
	РТ	0.5527	<u>0.0237</u>			
	TPB	<u>0.0483</u>	0.8841	<u>0.0015</u>		
r	AR					
	PM	0.8741				
	РТ	0.8810	0.8109			
	TPB	<u>0.0431</u>	<u>0.0343</u>	<u>0.0029</u>		

S3a. Post hoc statistical analysis P values among the levels of the ion exchanger.

Dognongo	Plasticizer (P value)						
Response		DOP	NPOE	DBP	NPPE	TCP	
0	DOP						
	NPOE	0.3347					
lop	DBP	<u>0.0322</u>	<u><0.0001</u>				
\mathbf{N}	NPPE	0.5144	0.6126	<u>0.0004</u>			
	ТСР	<u><0.0001</u>	<u><0.0001</u>	<u>0.0091</u>	<u><0.0001</u>		
	DOP						
\sim	NPOE	0.9152					
TOC	DBP	<0.0001	<0.001				
	NPPE	0.9211	0.9741	<0.0001			
	ТСР	0.9887	0.9847	<0.0001	0.9844		
ч	DOP						
	NPOE	<u><0.0001</u>					
	DBP	0.0518	<u><0.0001</u>				
	NPPE	0.9120	<u><0.0001</u>	0.1019			
	ТСР	0.9434	<u><0.0001</u>	<u>0.0371</u>	0.9270		

Table S3b. Post hoc statistical analysis *P* values among the levels of the plasticizer.

Dagnanga	Ionophore (P value)					
Response		CX-4	CX-6	CX-8	β-CD	β-CD/CX-8
Slope	CX-4					
	CX-6	<u><0.0001</u>				
	CX-8	<0.0001	0.6071			
	β-CD	<u><0.0001</u>	0.6275	0.9118		
	β-CD/CX-8	<0.0001	0.8379	0.8818	0.7825	
	CX-4					
\sim	CX-6	<u><0.0001</u>				
TOC	CX-8	<u><0.0001</u>	0.9795			
	β-CD	<u><0.0001</u>	0.9804	0.9812		
	β-CD/CX-8	<0.0001	0.9734	0.9776	0.9800	
ч	CX-4					
	CX-6	0.9183				
	CX-8	<u><0.0001</u>	<u><0.0001</u>			
	β-CD	0.1073	0.0847	<u>0.0271</u>		
	β-CD/CX-8	<u><0.0001</u>	<u><0.0001</u>	0.7414	<u>0.0440</u>	

Table S3c. Post hoc statistical analysis P values among the levels of the ionophore.

Domondo -	Conditioning solution Concentration (P value)				
Response		0.01	0.001	0.0001	
Slope	0.01				
	0.001	<u><0.0001</u>			
	0.0001	<u><0.0001</u>	<u><0.0001</u>		
LOQ	0.01				
	0.001	<u><0.0001</u>			
	0.0001	<u><0.0001</u>	<u><0.0001</u>		
T	0.01				
	0.001	<u><0.0001</u>			
	0.0001	<u><0.0001</u>	<u><0.0001</u>		

Table S3d. Post hoc statistical analysis P values among the levels of the conditioning solution concentration.

Dagnanga	Membrane Thickness (P value)					
Response		0.12	0.1	0.08		
lope	0.12					
	0.1	<u><0.0001</u>				
\sim	0.08	<u><0.0001</u>	<u><0.0001</u>			
LOQ	0.12					
	0.1	<u><0.0001</u>				
	0.08	<u><0.0001</u>	<u><0.0001</u>			
5	0.12					
	0.1	<u><0.0001</u>				
	0.08	<0.0001	<0.0001			

Table S3e. Post hoc statistical analysis *P* values among the levels of the membrane thickness.