Supplementary Table S1: Pharmacokinetics of the Components of DEC-C<sup>a</sup> (12)

Parameter	Decitabine	Cedazuridine
General Information		
	T	
Dose proportionality	20 mg to 40 mg with cedazuridine (100 mg)	40 to 100 mg with decitabine (20 mg)
With the recommended dosage of DE	C-C for 5 consecutive days:	
5-day cumulative AUC, ng·hr/mL	851 (50%)	
Day 1 AUC, ng·hr/mL	103 (55%)	2950 (49%)
Steady state AUC, ng·hr/mL	178 (53%)	3291 (45%)
Time to steady state, days	2	2
Accumulation ratio based on AUC	1.7 (42%)	1.1 (63%)
C <sub>max</sub> , ng/mL	145 (55%)	371 (52%)
Absorption		1
Bioavailability	Cedazuridine increases oral decitabine exposure	20% (23%)
<u>Elimination</u>		
Half-life at steady state, b hours	1.5 (27%)	6.7 (19%)
Metabolism		
Primary Pathways	Primarily by cytidine deaminase (CDA) and by physicochemical degradation	Conversion to epimer by physicochemical degradation
Excretion (healthy subjects)		
Total (% unchanged)		46% (21%) in urine and 51% (27%) in feces
Drug-Drug Interaction		
	Cedazuridine increases oral decitabine exposure	No effect of decitabine on cedazuridine
	DEC-C may increase systemic exposure of drugs metabolized by CDA.	
	• Proton pump inhibitors had no clinically meaningful effect on the exposures of DEC-C	

<sup>&</sup>lt;sup>a</sup> Mean (% coefficient of variation)

Abbreviations: AUC<sub>0-24h</sub>=area under the plasma concentration-time curve from time zero to 24 hours;  $C_{max}$ , maximum plasma concentration;  $T_{max}$ , time to maximum concentration; V/F, apparent volume of distribution; CL/F, apparent clearance.

<sup>&</sup>lt;sup>b</sup> Mean (standard deviation)