

Supplementary Tables

Supplementary Table S1. List of mitocans and non-mitocans used for bioinformatics analysis.

<i>Mitocans</i>	<i>Non-mitocans</i>
Cytarabine	Hydroxyurea
Doxorubicin	Fludarabine
Paclitaxel	BN-2629
Etoposide	E-7820
Rapamycin	Fulvestrant
Vinorelbine	Dasatinib
Temsirolimus	Pazopanib
Arsenic trioxide	Afatinib
Obatoclax	Dolastatin 10
Salinomycin	Estramustine
Staurosporine	Midostaurin
Simvastatin	Testolactone
Mitoxanthrone	Valrubicin
L-Asparaginase	Fostamatinib

Supplementary Table S2. List of cell lines included in the study.

Cell line	Cell line origin	Disease	Age	Sex	Prior treatment	Ploidy	Genetic mutations / rearrangements
MV-4-11	Peripheral blood	Acute myeloid leukemia	10	M	NA	2n=48 (hyperdiploid)	FLT3-ITD ¹ , t(4;11) ² [PMID: 12529668]
OCI-AML2	Peripheral blood	Acute myeloid leukemia	65	M	NA	Hyperdiploid 48(43-49)	<i>DNMT3A</i> [PMID: 21904384]
THP-1	Peripheral blood	Acute myeloid leukemia	1	M	NA	Near-diploid (46, XY)	<i>MLL-AF9</i> fusion, deletions in <i>CDKN2A</i> , <i>PTEN</i> [PMID: 11066077, 19635138]
MOLM-13	Peripheral blood	Acute myeloid leukemia	20	M	NA	Hyperdiploid 51(48-52)	FLT3-ITD, CBL deltaExon8 mutant, <i>MLL-AF9</i> fusion [PMID: 12529668; 9305600]
U251	Central nervous system	Glioblastoma	75	M	NA	2n=46	<i>CDKN2A</i> , <i>C9orf53</i> , <i>PTEN</i> , <i>TP53</i> [PMID: 24810477; 17088437]
SKOV3	Ovary	Adeno-carcinoma	64	F	Thiotepa	Hypodiploid (42-45)	<i>TP53</i> , <i>CDKN2A</i> , <i>PIK3CA</i> , <i>APC</i> [PMID: 26585234; 17088437]
OVCAR3	Ovary	Adeno-carcinoma	60	F	CyPh/CsPt/Adr	Aneuploid, near-triploid 69+/-	<i>TP53</i> [PMID: 17088437]

¹ internal tandem duplication

² chromosomal translocation

Supplementary Table S3. Basic characteristics of the patients, enrolled in the study.

N	PB/BM ¹	age	sex	WBC	% blast	Pretreatment	Mutation status: gene mutations/genetic rearrangements	Karyotype
1.	PB	29	F	4.6	36	Decitabine, Venetoclax, Dasatinib	PTPN11, CSF3R	48,XX,-11,del(17)(p12),+r,+2mar[2]/49,idem,+8[12]/48~50,XX,t(7;11)(q22;q24),+8,del(9)(q13q22),-11,del(17)(p12),+r,+2~3mar[cp6]
2.	PB	83	M	15.6	30	ONC201, Cytarabine	TERT, CSF3R, TP53, GATA2	45,XY,t(3;8)(q25;q22),-5,del(7)(q22q34),add(8)(q24.1),del(12)(p11.2),16,+mar[18]/46~49,idem,+del(7)(q22q34)[cp2]
3.	PB	35	F	23.2	62	Decitabine	ASXL1, ASXL2, NRAS, RUNX1, FLT3 ITD, PHF6	46, XX
4.	PB	53	M	2.1	63	DCLL9718S	KMT2A, U2AF1	46,XY,add(1)(p36.1),del(7)(q22q34),del(20)(q11.2q13.3)[19]/47,idem,+del(7)(q22q34)[1]
5.	PB	41	F	13.5	92	Hydroxyurea, Crenolanib, Decitabine, Venetoclax, Gilteritinib	TET2, FLT3 ITD	49,XX,+4,t(6;9)(p22;q34),+8,+15,del(15)(q11.2q21)x2[14]/48~50,idem,+13[cp6]
6.	PB	77	M	4.9	57	Azacitidine, Cytarabine, Cladribine, Venetoclax	TET2, SRSF2, RUNX1, CBL, IL7R	46, XY
7.	PB	32	F	12.7	5.5	No treatment	FLT3 ITD, NPM1, DNMT3A	46, XX
8.	PB	57	M	37.2	15	Hydroxyurea. Cytarabine	ASXL1, NRAS, TET2, RUNX1T1	47,XY,+19[1]/47,idem,der(22)t(8;22)(q21.2;p11.2)[11]/46,XY[8]
9.	PB	79	F	5.4	6	IMGN622	ASXL, BCOR, DNMT3A	46,XX,t(2;17)(p14;q12),add(10)(q24),del(21)(q22q22)[19]/46,XX[1]
10.	BM	80	M	128	89	Hydroxyurea	FLT3 ITD, RUNX1 ISH2, PIGA, DNMT3A, ZRSR2	46, XY
11.	BM	57	M	9.3	86	No treatment	SMC1A, SRSF2, IDH1, ASXL2, FLT3 ITD	47,XY,+8[3]/46,XY,add(5)(q35)[2]/48,XY,+8,+mar[7]/47,XY,add(5)(q35),+8[6]/48,XY,add(5)(q35),+8,+mar[1]/49,XY,+8,+2mar[1]
12.	PB	64	M	3.1	67	Decitabine	TET2, TP53, EZH2, CBL	41~45,Y,der(X)t(X;1)(q24;p13),-2,add(3)(p13),del(5)(q13q35),-8,psudic(9;10)(p22;p11.2),-15,-17,add(17)(p11.2),add(21)(p11.2),del(22)(q11.2),+2~6mar[cp19]/46,XY[1]
13.	PB	58	M	38	31	No treatment	ETV6, EZH2, NRAS, SETBP1, DNMT3A	46, XY
14.	PB	70	M	8.2	48	No treatment	KDM6A, RUNX1, TP53	46~47,XY,+1,der(1;10)(q10;q10),der(1;12)(p10;q10),del(5)(q11.2),del(6)(q15q25),-14,-16,add(16)(p11.2),-17,-21,+3~6mar[cp15]/46,XY[4]
15.	BM	89	M	95.8	65	No treatment	ASXL1, CBL, CREBBP, PHF6, TET2	50,XY,+der(1;11)(q10;q10),+8,+8,+8[10]/47~51,XY,+add(1)(p13),+8,+8,+8,der(9)t(1;9)(q21;p24)[cp10]
16.	BM	58	M	38	31	No treatment	ETV6, EZH2, NRAS, SETBP1, DNMT3A	46, XY
17.	PB	64	M	9.2	23	Dasatinib, Decitabine, Venetoclax	ASXL1	46,XY,add(2)(q12),t(4;12)(q12;q24.1),ins(12;?)(p13;?)[18]/46,XY[2]

18.	PB	72	M	5.2	90	AMG 330, Tocilizumab, Hydrea	BCOR, U2AF1, DNMT3A	46, XY
19.	PB	21	M	4.8	86	Decitabine, Venetoclax,	NPM1, IDH2	46,XY,t(2;4)(p23;q21)[3]/46,XY,t(5;17)(q13;q25) [2]/46,XY[15]
20.	PB	30	M	2.3	61	Mitoxantrone, Etoposide, Cytarabine	BCOR, PTPN11	45,XX,t(3;3)(q21;q26.2),del(6)(q12),-7[16]
21.	PB	72	M	1.4	38	Hydroxyurea, Quizartinib, Decitabine	CEBPA, FLT3 ITD, KRAS, PTPN11, WT1	46, XY

¹ Source of primary AML cells: PB (peripheral blood) / BM (bone marrow).

Supplementary Table S4. Primer sequences used for qPCR studies.

Locus	Forward (5'-3')	Reverse (5'-3')	PCR protocol
<i>Primers used for estimation of relative mtDNA content (mtDNA/gDNA ratio)</i>			I. 95 °C for 30 s; II. 1. 95 °C for 10 s; 2. 55 °C for 45 s; +Plate read (40 cycles)
<i>GAPDH</i>	TCTGACTTCAACAGCGACAC	CAGCCACATACCAGGAAATG	
<i>B2M</i>	TGTAAGCAGCATCATGGAGG	CTCCACATAGTGAGGGTTATC	
<i>TBP</i>	GAGAGTTCTGGGATTGTACC	CCATCTGAAAACAGAGCAGG	
<i>ACTB</i>	AAAGACCTGTACGCCAACAC	GGAAAGACACCCACCTTGAT	
<i>ND1</i>	GTATTATACCCACACCCACC	GTACAATGAGGAGTAGGAGG	
<i>ND4</i>	CACATATGGCCTAGACTACG	GTGGGTGGTTGTGTTGATTC	
<i>Primers used for gene expression studies</i>			III. Melt Curve 65 to 95 °C, increment 0.5 °C for 5 s +Plate read END
<i>MFN1</i>	TCTCCGCCTTTAACTTCTCG	TTATGCTAAGTCTCCGCTCC	
<i>MFN2</i>	GCAGGACATGATAGATGGCT	ATAGTTGAGGGAGAAGCACTG	
<i>OPA1</i>	GTCAGTCAAATGGACCCTCA	ATCTGCTGAATCCTGCTTGG	
<i>DNML1</i>	GAACAGCGAGATTGTGAGGTT	TGCATTACTGCCTTTGGCAC	
<i>FIS1</i>	ATCCGTAAAGGCATCGTGCT	AGAAGACGTAATCCCGCTGT	
<i>MFF</i>	ACCTGACTGTTGTAGATGCAG	AGCACGTTCTTTGTTCTCCTC	
<i>MID49</i>	GTGAAGCGGTTTCATTGACAG	TTGAGCAGGCTCAGTTCCTT	
<i>MID51</i>	ATGTACTTGAGTGGCAGCCT	GGGCACAATGAGTTGGATGT	
<i>ACTB</i>	GATCAAGATCATTGCTCCTCTG	ATACTCCTGCTTGCTGATCCA	

Supplementary Table S5. Results of multiple linear regression: association of cancer type with the sensitivity to mitocan treatment.

Dependent variable ¹	B-coefficient	SE(B)	<i>p</i> -value
<i>After adjustment for age, gender and prior treatment of the cell line donor</i>			
Mitocan activity Z-scores	0.382	0.168	0.031
Non-mitocan activity Z-scores	0.346	0.171	0.054

¹ With cancer type as an independent predictor
 $p < 0.05$ are highlighted in bold

Supplementary Table S6. Fold changes (positive – increase; negative – decrease) of mitochondrial and bioenergetic parameters after doxorubicin treatment (DOX/untreated).

Cell line	mtDNA content	MTG fluorescence	JC-1, red/green	ATP content	Basal OCR	ATP-linked OCR	Proton leak	Spare capacity	Coupling efficiency	ECAR	WB (ATP syn β)	LD50 (DOX), μM
MV-4-11	-1.49			-19.71	-2.19	-2.84	-1.94	-1.10	-1.43	1.22		1.634
OCI-AML2	-1.44			-26.80	-2.02	-3.26	-1.28	-4.26	-2.22	-1.03		21.67
MOLM-13	-1.77	-32.03	-63.96	-9.57	-3.11	-6.29	-1.23	-5.95	-2.16	-1.10	1.118	26.75
THP-1	-1.40			-5.66	-5.32	-9.26	-3.30	-3.80	-2.36	-1.92		68.70
PBMC	1.40	-13.97	-4.71	-8.76	-1.55	-1.61	-1.41	-2.23	-1.20	-1.52	3.772	40.85
U251	5.51			-1.05								122.22
SKOV3	2.37	-29.19	-15.34	1.07	-1.28	-1.26	-1.03	-1.90	-1.07	1.09	1.002	110.82
OVCAR3	5.59			1.05								93.87

Cases with p<0.05 are in **bold**.

Supplementary Table S7. Fold change differences in expression of fusion and fission genes in DOX-treated (LD50, 8 h) cells compared to untreated cells.

Gene Cell line	Fusion			Fission				
	<i>MFN1</i>	<i>MFN2</i>	<i>OPAI</i>	<i>DNM1L</i>	<i>FIS1</i>	<i>MFF</i>	<i>MID49</i>	<i>MID51</i>
MV-4-11	0.83 (0.50-1.37)*	2.13 (1.34-3.39)	1.35 (0.84-2.18)	2.06 (1.26-3.36)	3.06 (1.88-4.99)	1.99 (1.19-3.34)	5.71 (3.52-9.27)	2.86 (1.69-4.85)
MOLM-13	1.09 (0.53-2.26)	2.19 (0.76-6.32)	0.51 (0.25-1.03)	0.58 (0.24-1.38)	0.80 (0.30-2.12)	0.53 (0.28-1.01)	16.82 (4.96-56.95)	1.41 (0.42-4.71)
THP-1	5.43 (3.71-7.95)	6.95 (3.61-13.37)	0.71 (0.52-0.98)	0.36 (0.13-0.99)	1.84 (1.26-2.69)	0.66 (0.48-0.92)	32.98 (15.96-68.15)	3.14 (1.78-5.54)
PBMC	3.09 (1.11-8.63)	4.82 (1.18-19.67)	1.29 (0.47-3.59)	0.91 (0.45-1.84)	1.52 (0.74-3.10)	2.24 (1.04-4.84)	13.67 (2.79-67.09)	4.72 (1.08-20.68)
SKOV3	0.33 (0.11-0.95)	0.67 (0.51-0.86)	0.46 (0.17-1.21)	0.52 (0.18-1.51)	0.93 (0.72-1.21)	0.63 (0.35-1.12)	0.58 (0.37-0.89)	0.25 (0.21-0.30)

* Fold changes were calculated using the comparative $\Delta\Delta C_t$ method

Supplementary Table S8. Relative potency of drugs used for combination studies (single-ray design) in cell lines and primary AML samples.

Cell line/Patient sample	LD25 _{1st drug} / LD25 _{2nd drug} = Relative potency	Fixed ratio (50:50) concentrations of the 1 st drug, 2 nd drug
<i>2-DG (1st drug) / CCCP (2nd drug)</i>		
MV-4-11	7500 μ M / 2.07 μ M = 3623	10.7 mM, 3 μ M; 7.5 mM, 2.1 μ M; 3.6 mM, 1 μ M
OCI-AML2	16240 μ M / 15.63 μ M = 1039	16.2 mM, 15.6 μ M; 8.1 mM, 7.8 μ M; 6.55 mM, 6.3 μ M
MOLM-13	51600 μ M / 6.3 μ M = 8190	51.6 mM, 6.3 μ M; 41 mM, 5 μ M.
THP-1	74800 μ M / 4.06 μ M = 18424	74.8 mM, 4.1 μ M; 36.8 mM, 2 μ M; 18.4 mM, 1 μ M
PBMC	106000 μ M / 45.41 μ M = 2334	106 mM, 45.41 μ M; 70 mM, 30 μ M; 46.7 mM, 20 μ M; 23.3 mM, 10 μ M
U251	616150 μ M / 92.3 μ M = 6676	616.15 mM, 92.3 μ M; 467.3 mM, 70 μ M; 333.8 mM, 50 μ M
SKOV3	737000 μ M / 133.4 μ M = 5525	800 mM, 139 μ M; 737 mM, 133.4 μ M; 688.9 mM, 120 μ M
OVCAR3	390748 μ M / 130.3 μ M = 2999	450 mM, 150 μ M; 390.7 mM, 130.3 μ M; 300 mM, 100 μ M; 150 mM, 50 μ M; 30 mM, 10 μ M
AML 1	8660 μ M / 12.33 μ M = 702	8.66 mM, 12.33 μ M; 5.77 mM, 8.22 μ M; 2.87 mM, 4.11 μ M
AML 2	79100 μ M / 16.07 μ M = 4922	79 mM, 16.07 μ M; 50 mM, 10 μ M; 24.5 mM, 5 μ M
AML 3	80700 μ M / 11.4 μ M = 7079	80.7 mM, 11.4 μ M; 53 mM, 7.5 μ M; 25 mM, 3.5 μ M
AML 4	66200 μ M / 7.7 μ M = 8597	66.2 mM, 7.7 μ M; 43 mM, 5 μ M; 21.5 mM, 2.5 μ M
AML 5	80900 μ M / 8.3 μ M = 9747	80.9 mM, 8.3 μ M; 57.8 mM, 6 μ M; 40.5 mM, 4 μ M; 19.5 mM, 2 μ M
AML 6	3700 μ M / 2.6 μ M = 1423	3.7 mM, 2.6 μ M; 2.9 mM, 2 μ M; 1.4 mM, 1 μ M
AML 7	23800 μ M / 17.4 μ M = 1368	23.8 mM, 17.4 μ M; 13.7 mM, 10 μ M; 6.8 mM, 5 μ M
AML 8	58900 μ M / 23.8 μ M = 2475	58.9 mM, 23.8 μ M; 36.8 mM, 15 μ M; 24.5 mM, 10 μ M; 12.3 mM, 5 μ M
AML 9	268000 μ M / 13.5 μ M = 19852	268 mM, 13.5 μ M; 198.5 mM, 10 μ M; 99.3 mM, 5 μ M; 19.9 mM, 1 μ M
AML 10	58800 μ M / 2.23 μ M = 26368	58.8 mM, 2.23 μ M; 26.4 mM, 1 μ M; 13.2 mM, 0.5 μ M
AML 11	4800 μ M / 14.5 μ M = 331	4.8 mM, 14.5 μ M; 2.4 mM, 7.25 μ M; 1.2 mM, 3.6 μ M
AML 12	56100 μ M / 10.33 μ M = 5431	56.1 mM, 10.3 μ M; 28.1 mM, 5.2 μ M; 14 mM, 2.6 μ M; 7 mM, 1.3 μ M
AML 13	200700 μ M / 55.1 μ M = 3642	200 mM, 55.1 μ M; 100 mM, 27.6 μ M; 50 mM, 13.8 μ M
AML 14	43200 μ M / 39.6 μ M = 1091	43.2 mM, 39.6 μ M; 21.6 mM, 19.8 μ M;

		10.8 mM, 9.9 μ M
AML 15	7950 μ M / 70.2 μ M = 113	8 mM, 70 μ M; 4 mM, 35 μ M; 2 mM, 17.5 μ M; 1 mM, 8.75 μ M
AML 16	95700 μ M / 17 μ M = 5629	95.7 mM, 17 μ M; 47.9 mM, 8.5 μ M; 23.9 mM, 4.25 μ M; 12 mM, 2.13 μ M
AML 17	81000 μ M / 16.7 μ M = 4850	81 mM, 16.7 μ M; 40.5 mM, 8.35 μ M; 20.25 mM, 4.18 μ M; 10.1 mM, 2.1 μ M
AML 18	10300 μ M / 4 μ M = 2575	10.3 mM, 4 μ M; 2.6 mM, 1 μ M; 1.3 mM, 0.5 μ M
AML 19	18200 μ M / 0.55 μ M = 33091	18.2 mM, 0.55 μ M; 12.1 mM, 0.37 μ M; 9.1 mM, 0.28 μ M; 4.55 mM, 0.13 μ M; 2.28 mM, 0.07 μ M
AML 20	850 μ M / 2.18 μ M = 390	0.85 mM, 2.2 μ M; 0.425 mM, 1.1 μ M; 0.2 mM, 0.55 μ M; 0.1 mM, 0.27 μ M
AML 21	105300 μ M / 22.8 μ M = 4618	105.3 mM, 22.8 μ M; 52.7 mM, 11.4 μ M; 26.3 mM, 5.7 μ M
<i>CCCCP (1st drug) / ABT-199 (2nd drug)</i>		
OCI-AML2	15.63 μ M / 0.177 μ M = 88	15.63 μ M, 0.177 μ M; 10 μ M, 0.113 μ M; 5 μ M, 0.06 μ M; 1 μ M, 0.0113 μ M
PBMC	45.41 μ M / 0.6 μ M = 76	45.1 μ M, 0.6 μ M; 30 μ M, 0.4 μ M; 20 μ M, 0.26 μ M; 10 μ M, 0.13 μ M
<i>2-DG (1st drug) / ABT-199 (2nd drug)</i>		
OCI-AML2	16240 μ M / 0.177 μ M = 91751	16.24 mM, 0.177 μ M; 13.8 mM, 0.15 μ M; 9.2 mM, 0.1 μ M; 1.8 mM, 0.02 μ M
PBMC	106000 μ M / 0.6 μ M = 176667	106 mM, 0.6 μ M; 53 mM, 0.3 μ M; 26.5 mM, 0.15 μ M; 10.6 mM, 0.06 μ M
<i>2-DG (1st drug) / ara-C (2nd drug)</i>		
OCI-AML2	16240 μ M / 77.8 μ M = 209	16.2 mM, 77.8 μ M; 8.1 mM, 38.9 μ M; 4.1 mM, 19.5 μ M; 2 mM, 9.7 μ M
AML 11 ¹	4800 μ M / 53.4 μ M = 90	4.8 mM, 53.4 μ M; 2.4 mM, 26.7 μ M; 1.2 mM, 13.4 μ M
AML 13	200700 μ M / 131.8 μ M = 1523	200 mM, 131.8 μ M; 100 mM, 69.1 μ M; 50 mM, 34.5 μ M
AML 14	43150 μ M / 400 μ M = 108	43.2 mM, 400 μ M; 21.6 mM, 200 μ M; 10.8 mM, 100 μ M
AML 15	7950 μ M / 50.4 μ M = 158	8 mM, 50 μ M; 4 mM, 25 μ M; 2 mM, 12.5 μ M; 1 mM, 6.25 μ M
AML 20	850 μ M / 141.2 μ M = 6	0.85 mM, 141.2 μ M; 0.425 mM, 70.6 μ M; 0.2 mM, 35.3 μ M; 0.1 mM, 17.7 μ M

¹ Only primary AML samples, not resistant to ara-C (LD25 < 400 μ M corresponding to 0.4% DMSO), have been treated with combination of ara-C and 2-DG.

Supplementary Table S9. Results of multiple linear regression: no association of pretreatment of the patients (n = 21) with mean combination indices of CCCP/2-DG.

Dependent variable ¹	B-coefficient	SE(B)	p-value
<i>Before adjustment for age, gender, source of primary cells and % blast cells</i>			
Mean combination index based on LD25, 50, 75 of the mixture CCCP/2DG	-0.085	0.118	0.481
<i>After adjustment for age, gender, source of primary cells and % blast cells</i>			
Mean combination index based on LD25, 50, 75 of the mixture CCCP/2DG	0.002	0.153	0.988

¹ With pretreatment (0 / 1) as an independent predictor

Supplementary Table S10. Correlations between mitochondrial parameters across untreated AML cells, healthy blood cells and SKOV3 cell line¹.

		Max OCR	ATP-linked OCR	Spare capacity	Coupling efficiency	LD25, CCCP	LD25, 2-DG	CI based on LD50 of the mixture	Mean CI based on LD25,50,75 of the mixture
Basal OCR	<i>r</i>	0.915	0.975	0.790	0.189	0.345	0.565	0.521	0.283
	<i>p</i>	0.011	0.001	0.061	0.719	0.500	0.243	0.289	0.587
Max OCR	<i>r</i>		0.966	0.969	0.477	0.463	0.644	0.579	0.397
	<i>p</i>		0.002	0.001	0.338	0.355	0.168	0.229	0.436
ATP-linked OCR	<i>r</i>			0.888	0.376	0.500	0.694	0.639	0.424
	<i>p</i>			0.018	0.463	0.312	0.126	0.172	0.402
Spare capacity	<i>r</i>				0.651	0.550	0.687	0.61	0.481
	<i>p</i>				0.161	0.258	0.132	0.199	0.335
Coupling efficiency	<i>r</i>					0.840	0.784	0.749	0.821
	<i>p</i>					0.036	0.065	0.087	0.045
LD25, CCCP	<i>r</i>						0.969	0.907	0.893
	<i>p</i>						0.001	0.012	0.017
LD25, 2-DG	<i>r</i>							0.943	0.869
	<i>p</i>							0.005	0.025
CI based on LD50 of the mixture	<i>r</i>								0.960
	<i>p</i>								0.002

¹ Correlations defined by two-tailed Pierson correlation coefficient *r*; *p* defines *p*-value. Significant correlations (*p*<0.05) are in bold.

Supplementary Table S11. Changes in mitochondrial bioenergetic parameters after treatment with mitocans (CCCP, DOX, LD50), glycolysis inhibitor (2-DG, LD50), or their combination (CCCP and 2-DG, LD25) in a panel of cells.

Drug	Bioenergetic parameter	MV-4-11	OCI-AML2	MOLM-13	THP-1	PBMC	SKOV3
CCCP, LD50	Basal OCR	$p < 0.0001$ -74%	$p < 0.0001$ -92%	$p < 0.0001$ -89%	$p < 0.0001$ -96%	NS	$p < 0.0001$ -79%
	Maximal OCR	$p < 0.0001$ -67%	$p < 0.0001$ -96%	$p < 0.0001$ -96%	$p < 0.0001$ -98%	$p < 0.0001$ -88%	$p < 0.0001$ -88%
	ATP-linked OCR	$p < 0.0001$ -101%	$p < 0.0001$ -97%	$p < 0.0001$ -96%	$p < 0.0001$ -95%	$p = 0.0020$ -71%	$p < 0.0001$ -100%
	Proton leak	NS	$p < 0.0001$ -85%	$p < 0.0001$ -68%	$p < 0.0001$ -100%	$p = 0.0034$ +147%	NS
	Spare capacity	$p = 0.0211$ -64%	$p < 0.0001$ -100%	$p < 0.0001$ -100%	$p < 0.0001$ -100%	$p < 0.0001$ -105%	$p < 0.0001$ -96%
	Coupling efficiency	$p < 0.0001$ -116%	$p = 0.0161$ -31%	$p = 0.0017$ -19%	$p < 0.0001$ -45%	$p = 0.0002$ -105%	$p < 0.0001$ -129%
	ECAR	NS	$p < 0.0047$ -19%	$p = 0.0001$ -49%	$p = 0.0004$ -57%	NS	NS
DOX, LD50	Basal OCR	$p < 0.0001$ -54%	$p = 0.0006$ -51%	$p < 0.0001$ -68%	$p = 0.0002$ -81%	$p = 0.0021$ -35%	NS
	Maximal OCR	$p = 0.0108$ -30%	$p = 0.0007$ -62%	$p < 0.0001$ -77%	$p < 0.0001$ -77%	$p = 0.0097$ -52%	NS
	ATP-linked OCR	$p < 0.0001$ -65%	$p = 0.0012$ -69%	$p = 0.0002$ -84%	$p = 0.0010$ -89%	$p = 0.0340$ -38%	NS
	Proton leak	NS	NS	NS	$p = 0.0090$ -70%	NS	NS
	Spare capacity	NS	$p = 0.0051$ -77%	$p = 0.0034$ -83%	$p = 0.0074$ -74%	$p = 0.0194$ -55%	$p = 0.0175$ -47%
	Coupling efficiency	$p = 0.0062$ -30%	$p = 0.0119$ -55%	NS	$p = 0.0157$ -58%	NS	NS
	ECAR	NS	NS	NS	$p = 0.0087$ -48%	$p = 0.0020$ -34%	NS
2-DG, LD50	Basal OCR	$p < 0.0001$ -62%	$p = 0.0047$ -38%	$p < 0.0001$ -52%	$p = 0.0020$ -76%	NS	$p < 0.0001$ -47%
	Maximal OCR	NS	NS	$p = 0.0017$ -27%	$p < 0.0001$ -53%	NS	$p = 0.0242$ -52%
	ATP-linked OCR	$p < 0.0001$ -63%	$p = 0.0270$ -50%	NS	NS	NS	$p < 0.0001$ -71%
	Proton leak	$p = 0.0076$ -65%	NS	$p = 0.0188$ -35%	$p = 0.0068$ -63%	NS	$p = 0.0004$ +97%
	Spare capacity	$p = 0.0017$ +86%	NS	NS	NS	NS	NS
	Coupling efficiency	NS	NS	NS	NS	NS	$p < 0.0001$ -57%
	ECAR	NS	$p < 0.0001$ -37%	$p = 0.0017$ -47%	$p = 0.0014$ -56%	$p < 0.0001$ -60%	NS
CCCP, LD25 + 2-DG, LD25	Basal OCR	$p < 0.0001$ -81%	$p = 0.0066$ -42%	$p < 0.0001$ -83%	NS	$p < 0.0001$ -70%	$p < 0.0001$ -95%
	Maximal OCR	$p < 0.0001$ -52%	$p < 0.0001$ -72%	$p < 0.0001$ -89%	$p < 0.0001$ -61%	$p < 0.0001$ -94%	$p < 0.0001$ -96%
	ATP-linked OCR	$p < 0.0001$ -95%	$p < 0.0001$ -88%	$p < 0.0001$ -96%	$p = 0.0103$ -87%	$p < 0.0001$ -100%	$p < 0.0001$ -101%

	Proton leak	$p = 0.0056$ -67%	NS	$p = 0.0038$ -42%	NS	NS	NS
	Spare capacity	NS	$p < 0.0001$ -107%	$p < 0.0001$ -92%	NS	$p < 0.0001$ -99%	$p < 0.0001$ -98%
	Coupling efficiency	$p = 0.0003$ -72%	$p < 0.0001$ -80%	$p < 0.0001$ -67%	$p = 0.0005$ -64%	$p = 0.0016$ -85%	$p < 0.0001$ -121%
	ECAR	NS	NS	$p < 0.0001$ -58%	NS	$p < 0.0001$ -88%	$p = 0.0095$ -41%

ANOVA with subsequent Dunn's or Fisher LSD test was used. Significant changes ($p < 0.05$) are in bold.