



**Figure 1.- CONSORT flow diagram.** Patients who discontinued medication were not excluded from the final analysis.

	<b>Sulodexide n= 124</b>	<b>Placebo n= 119</b>	<b>Relative Risk (95%CI)</b>	<b>P-value</b>
<b>Demographics</b>				
Age in years, mean (SD)	55.3 (10.3)	54 (10.9)	..	0.26
Gender, n (%)				
Male	60 (48.3)	55 (46.2)	1.04 (0.80 - 1.36)	0.73
Female	64 (51.6)	64 (53.8)	0.95 (0.75 - 1.21)	0.73
BMI, Mean (SD)	29 (4.0)	28.7 (3.2)	..	0.30
<b>Chronic comorbidities, n (%)</b>				
Diabetes mellitus	22 (17.7)	28 (23.5)	0.75 (0.45 - 1.24)	0.26
Hypertension	48 (38.7)	35 (29.1)	1.31 (0.92 - 1.87)	0.13
COPD	30 (24.1)	26 (21.8)	1.10 (0.69 - 1.75)	0.66
Cardiovascular disease	28 (22.5)	23 (19.32)	1.16 (0.71 - 1.90)	0.53
*C19HC risk calculator, mean (SD)	67.87 (14.0)	65.81 (14.1)	..	0.32

Table 1. Results Demographic characteristics. \* *Percentage is given by the COVID-19 Health Complication (C19HC) risk calculator (Gobierno de Mexico, IMSS).*

<b>Outcome measures</b>				
Need Hospital care, n (%)	22 (17.7)	35 (29.4)	0.60 (0.37 - 0.96)	0.03
LOD hospital care, mean (SD)	6.29 (4.1)	7.8 (4.5)	..	0.21
†Need Oxygen support, n (%)	37 (29.8)	50 (42.0)	0.71 (0.50 - 1.00)	0.05
†LOD Oxygen support, mean (SD)	9 (7.2)	11.5 (9.6)	..	0.02
Mortality	3 (2.4)	7 (5.8)	0.41 (0.10 - 1.55)	0.19
Invasive Mechanical ventilation support, n (%)	3 (2.4)	6 (5.0)	0.47 (0.12 - 1.87)	0.29
Haemodialysis, n (%)	0	0	..	..
Thromboembolic events, n (%)	2 (1.6)	2 (1.6)	0.95 (0.13 - 6.70)	0.96
<b>Laboratory findings</b>				
D-dimer, ng/dl				
Baseline, mean (SD)	293.6 (117.5)	318.4 (131.6)	..	0.12
< 500, n (%)	110 (88.7)	98 (75.6)	1.07 (0.97 - 1.19)	0.16

> 500, n (%)	14 (11.2)	21 (17.6)	0.63 (0.34 - 1.19)	0.16
Week 2, mean (SD)	464.75 (629.81)	897.76 (1215.36)	..	<0.01
< 500, n (%)	97 (78.3)	63 (52.95)	1.47 (1.21 - 1.79)	<0.01
> 500, n (%)	27 (21.7)	56 (47.05)	0.46 (0.31 - 0.67)	<0.01
CRP, mg/dl				
Baseline, mean (SD)	10.6 (6.4)	10.1 (6.9)	..	0.55
Week 2, mean (SD)	12.55 (10.2)	17.81 (11.56)	..	<0.01
Creatinine week 2, mg/dl				
< 1.6	113 (91.1)	107 (89.9)	1.01 (0.93 - 1.09)	0.74
> 1.6	11 (08.8)	12 (10.0)	0.87 (0.40 - 1.91)	0.74

**Table 2. Results outcome**

† Includes the total of days that patients needed oxygen support at home or the hospital. Some patients continued oxygen support at-home after hospital care or started oxygen at home and later required hospital care.

LOD = length of days, SD = standard deviation, n = number patients, % = percentage, BMI = body mass index.

COPD = Chronic obstructive pulmonary disease. CRP = C-reactive protein.

	<b>Sulodexide N=124</b>	<b>Placebo N=119</b>	<b>Relative risk (CI 95%)</b>	<b>P-value</b>
<b>Medication adherence, n, (%)</b>				
†All the time	91 (73.3)	99 (83.1)	1.13 (0.99 - 1.29)	0.06
†Most of the time	14 (11.2)	8 (6.7)	0.59 (0.25 - 1.36)	0.22
Suspended medication				
total	19 (15.3)	12 (10)	1.81 (0.88 - 3.74)	0.10
*Adverse event	8 (6.4)	6 (5)	0.78 (0.27 - 2.18)	0.63
** Voluntary	11 (8.8)	6 (5)	0.56 (0.21 - 1.48)	0.24
<b>Adverse event, n, (%)</b>				
‡Total	96 (77.4)	85 (71.4)	1.08 (0.93 - 1.25)	0.28
Abdominal discomfort (gastritis, nausea, vomiting or diarrhoea).	36 (29)	39 (32.7)	1.12 (0.77 - 1.64)	0.52

Headache.	96 (77.4)	85 (71.4)	0.92 (0.79 - 1.07)	0.28
Major Bleeding.	0	1 (0.8)	3.12 (0.12 - 75.96)	0.48
Skin reaction.	3 (2.41)	5 (4.2)	1.73 (0.42 - 7.10)	0.44

**Table 3.- Medication adherence and adverse events.** Values are through day-21, the date of the scheduled completion of the trial intervention.

† Patients while on the per-protocol outpatient setting.

The main reason for the voluntary suspension of medication was symptoms improvement.

\* Three patients in the control group and five patients in the study group that suspended medication due to an adverse event required hospital care due to severe clinical disease progression.

\*\* No patient that suspended medication for voluntary reasons needed hospital care or oxygen support.

‡ More than one adverse event could occur per patient.

CI= confidence interval.