




## Delta LSM (kPa/year)

Grey: $\leq-1$ ( 0 death $/ 36$ )
Yellow: $-1<$ et <1 (5 deaths /162)

Orange: $\geq 1$ ( 6 deaths /48)


Delta LSM $\leq 0$ (8 deaths /61)

Delta LSM > 0 (15 deaths /42)

| Delta LSM $\leq 0$ | 61 | 60 | 52 | 40 | 22 | 7 |
| :--- | :--- | :--- | :--- | :--- | :--- | :--- |
| Delta LSM >0 | 42 | 38 | 34 | 25 | 13 | 2 |



Green : Baseline FIB-4 <1.12 : 6 deaths/425
Orange : $1.12 \leq$ Baseline FIB-4 <2.25: 6 deaths/378
Red: 2.25 S Baseline FIB-4 < 5.8 : 13 deaths/177

Black: Baseline FIB-4 $\geq 5.8: 17$ deaths/44


## Delta FIB-4 (unit/year)

Green: <0.20:17 deaths/818
Orange : $\geq 0.20$ and $<1.40$ : 13 deaths/182

Red: $\geq 1.40: 12$ deaths/23

| Green | 818 | 811 | 749 | 591 | 355 | 93 |
| :--- | :---: | :---: | :---: | :---: | :---: | :---: |
| Orange | 182 | 178 | 162 | 121 | 74 | 27 |
| Red | 23 | 20 | 18 | 13 | 8 | 0 |

## Baseline LSM



Green: <7.0 kPa (0 death /574)
Orange: $7.0-13.9 \mathrm{kPa}$ (7 deaths /321)
Red: $\geq 14.0 \mathrm{kPa}$ ( 14 deaths $/ 130$ )

| Green | 574 | 567 | 518 | 403 | 243 | 66 |
| :--- | :---: | :---: | :---: | :---: | :---: | :---: |
| Orange | 321 | 319 | 300 | 237 | 147 | 39 |
| Red | 130 | 125 | 113 | 85 | 47 | 15 |

Panel s9a


| Blue | 574 | 567 | 518 | 403 | 243 | 56 |
| :--- | :---: | :---: | :---: | :---: | :---: | :---: |
| Green | 102 | 102 | 98 | 75 | 47 | 14 |
| Grey | 36 | 36 | 35 | 24 | 16 | 6 |
| Kellow | 162 | 162 | 153 | 126 | 80 | 19 |
| Orange | 48 | 46 | 41 | 32 | 15 | 6 |
| Red | 61 | 60 | 52 | 40 | 22 | 7 |
| Black | 48 | 38 | 34 | 25 | 13 | 2 |


| Group | Baseline LSM <br> $(\mathrm{hPa})$ | Delta LSM <br> $(\mathrm{kPa} /$ year) | SVR |
| :--- | :--- | :--- | :--- |
| 1 (Blue) | $<7$ |  |  |
| 2 (Green) | 27 |  | Yes |
| 3 (Grev) | 27 and $<14$ | $\$ 1$ | No |
| 4 (Yellow) | 27 and $<14$ | $>-1$ and $<1$ | No |
| 5 (Orange) | 27 and $<14$ | $\geq 1$ | No |
| 6 (Red) | 214 | $\leq 0$ | No |
| 7 (Black) | 214 | $>0$ | No |

## Panel $59 b$



| Violet | 425 | 424 | 395 | 308 | 196 | 53 |
| :--- | :---: | :---: | :---: | :---: | :---: | :---: |
| Blue | 303 | 298 | 274 | 220 | 134 | 38 |
| Green | 75 | 72 | 66 | 50 | 33 | 11 |
| Grey | 134 | 133 | 123 | 95 | 57 | 11 |
| Yellow | 34 | 33 | 30 | 23 | 15 | 6 |
| Orange | 9 | 7 | 7 | 5 | 2 | 0 |
| Bed | 23 | 23 | 19 | 15 | 4 | 1 |
| Brown | 8 | 8 | 5 | 2 | 1 | 0 |
| Black | 13 | 12 | 10 | 7 | 5 | 0 |


| Group | Baseline FIB-4 | Delta FIB-4 (unit/year) |
| :--- | :--- | :--- |
| $\mathbf{1}$ (Violet) | $<1.12$ |  |
| $\mathbf{2}$ (Blue) | $\geq 1.12$ and $<2.25$ | $<0.20$ |
| 3 (Green) | $\geq 1.12$ and $<2.25$ | $\geq 0.20$ |
| $\mathbf{4}$ (Grey) | $\geq 2.25$ and $<5.80$ | $<0.20$ |
| $\mathbf{5}$ (Yellow) | $\geq 2.25$ and $<5.80$ | $\geq 0.20$ and $<1.40$ |
| $\mathbf{6}$ (Orange) | $\geq 2.25$ and $<5.80$ | $\geq 1.40$ |
| 7 (Red) | $\geq 5.80$ | $<0.20$ |
| $\mathbf{8}$ (Brown) | $\geq 5.80$ | $\geq 0.20$ and $<1.40$ |
| 9 (Black) | $\geq 5.80$ | $\geq 1.40$ |

Supplementary table s1: Comparison between baseline and follow-up results of fibrosis tests as a function of treatment response.

| Fibrosis test | Treatment <br> response | Baseline | Follow-up | p | Rs (p) | Delta <br> (kPa or unit/year) |
| :--- | :--- | :--- | :--- | :--- | :--- | :--- |
| Fibroscan | All | $6.6(5.1-9.5)$ | $6.2(4.8-8.8)$ | 0.014 | $0.650(<0.001)$ | $-0.07(-0.55-0.38)$ |
| $\mathbf{( k P a )}$ | No SVR | $6.3(5.0-9.0)$ | $6.1(4.8-9.0)$ | 0.756 | $0.681(<0.001)$ | $-0.03(-0.45-0.40)$ |
|  | SVR | $8.1(5.7-11.8)$ | $6.3(4.8-8.5)$ | $<0.001$ | $0.522(<0.001)$ | $-0.40(-1.25-0.18)$ |
| FIB-4 | All | $1.27(0.84-2.03)$ | $1.49(1.04-2.30)$ | $<0.001$ | $0.742(<0.001)$ | $0.05(-0.02-0.16)$ |
|  | No SVR | $1.25(0.83-1.95)$ | $1.51(1.05-2.37)$ | $<0.001$ | $0.776(<0.001)$ | $0.06(0.00-0.16)$ |
|  | SVR | $1.45(0.87-2.31)$ | $1.42(0.94-2.12)$ | 0.628 | $0.587(<0.001)$ | $0.01(-0.14-0.12)$ |
| APRI | All | $0.43(0.27-0.77)$ | $0.42(0.27-0.76)$ | 0.002 | $0.569(<0.001)$ | $0.01(-0.04-0.05)$ |
|  | No SVR | $0.41(0.27-0.70)$ | $0.45(0.31-0.81)$ | $<0.001$ | $0.666(<0.001)$ | $0.01(-0.02-0.06)$ |
|  | SVR | $0.56(0.29-1.09)$ | $0.27(0.19-0.48)$ | $<0.001$ | $0.434(<0.001)$ | $-0.05(-0.20-0.01)$ |

Results are expressed as median with 1st and 3rd quartiles into brackets. Rs: Spearman correlation coefficient; SVR: sustained viral response

Supplementary table s2: Rate of death as a function of patient subgroups defined by baseline and delta LSM results.

| Baseline LSM <br> (kPa) | Delta LSM <br> (kPa/year) |  | SVR | Patients | Rate (\%) of death during follow-up |  |
| :--- | :--- | :--- | :--- | :--- | :--- | :---: |
|  | - | - | 574 | 1.0 | 0.0 |  |
| $<7.0$ | - | Yes | 102 | 2.0 | 1.0 |  |
| $\geq 7.0$ | - | No | 36 | 0.0 | 0.0 |  |
| $\geq 7.0$ and $<14.0$ | $\leq-1.0$ | No | 162 | 3.1 | 1.2 |  |
|  | $-1.0<$ and $<1.0$ | No | Liver-related |  |  |  |
|  | $\geq 1.0$ | No | 48 | 12.5 | 10.4 |  |
| $\geq 14.0$ | $\leq 0$ | No | 61 | 13.1 | 6.6 |  |
|  | $>0$ | No | 42 | 35.7 | 21.4 |  |

SVR: sustained viral response

Supplementary table s3: Rate of death as a function of patient subgroups defined by baseline and delta FIB-4 results.

| Baseline FIB-4 | Delta FIB-4 <br> (unit/year) | Patients <br> (n) | Rate (\%) of death during follow-up |  |
| :---: | :---: | :---: | :---: | :---: |
|  |  |  | Overall | Liver-related |
| $<1.12$ | - | 425 | 1.4 | 0.2 |
| $\geq 1.12$ and $<2.25$ | <0.20 | 303 | 1.0 | 0.3 |
|  | $\geq 0.20$ | 75 | 4.0 | 1.3 |
| $\geq 2.25$ and $<5.8$ | <0.20 | 134 | 3.7 | 1.5 |
|  | $\geq 0.20$ and $<1.40$ | 34 | 8.8 | 2.9 |
|  | $\geq 1.40$ | 9 | 55.6 | 44.4 |
| $\geq 5.8$ | $<0.20$ | 23 | 21.7 | 17.4 |
|  | $\geq 0.20$ and $<1.40$ | 8 | 62.5 | 25.0 |
|  | $\geq 1.40$ | 13 | 53.8 | 38.4 |

