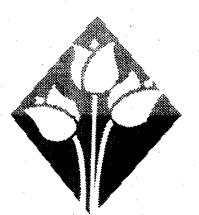
## RESULTS



## Results

The results of the present study were summarized, statistically analyzed and presented in (11) tables and (10) figures.

Table (12): Age distribution among the study population.

| Age (Year) | Patients (N=21) | Control (N=20) |  |  |  |
|------------|-----------------|----------------|--|--|--|
| Mean ± SD  | 47.43 ± 10.67   | 49.2 ± 8.54    |  |  |  |
| T value    | 0.588           |                |  |  |  |
| P value    | 0.560           |                |  |  |  |

The mean age of the patient group was  $47.43. \pm 10.67$  years while, the mean age of the control group was  $49.22 \pm 8.54$  years. The analysis of difference between the age in patients and control groups was statistically non significant (P > 0.05).

Table (13): Sex distribution among the study population.

| Sex     | Patients (N=21) | Controls (N=20) |  |
|---------|-----------------|-----------------|--|
| Males   | 17 (80.95%)     | 12 (60%)        |  |
| Females | 4 (19.05%)      | 8 (40%)         |  |

Pearson Chi - Square

Value = 2.17

df=1

P = 0.141

The patients group included 21 patients with chronic hepatitis C 17 males (80.95%) and 4 females (19.05%). The control group included 20 HCV - negative individual 12 males 12 (60%) and 8 females 8 (40%). The analysis of difference between the sex among patients and control groups was statistically non significant (P > 0.05).

Table (14) Analysis of difference between surface markers of the peripheral blood lymphocytes (PBL) in patients and control group (unpaired t- test):

| Surface marker<br>(%)                     | Patients (N=21) Mean ± SD | Control (N=20) Mean ± SD | t value | P value |
|---|---------------------------|--------------------------|---------|---------|
| - T – cell                                |                           |                          |         |         |
| CD3 <sup>+</sup>                          | 65.6 ± 9.52               | 60.9 ± 4.8               | 2.046   | 0.051   |
| CD4 <sup>+</sup>                          | 44.1 ± 7.32               | 35.7 ± 3.34              | 4.664   | 0.001   |
| CD8 <sup>+</sup>                          | 22.1 ± 8.32               | 25.2 ± 3.7               | -1.583  | 0.125   |
| CD4 <sup>+</sup> / CD8 <sup>+</sup> ratio | 2.39 ± 1.1                | 1.45 ± 0.27              | 3.776   | 0.001   |
| - B – cell (CD19 <sup>+</sup> )           | 6.17 ± 3.14               | 5.97 ± 2.26              | 1.42    | 0.21    |
| - NK (CD56 <sup>+</sup> )                 | 14.02 ± 5.04              | 12.65 ± 3.59             | 1.07    | 0.316   |

This table shows the mean ± SD of the surface markers of the PBL in patients and control groups. There was a statistically significant difference between the proportion of the peripheral blood CD4<sup>+</sup> and CD4<sup>+</sup> / CD8<sup>+</sup> ratio among the patients and the control groups. There was no statistically significant difference between the proportion of CD3<sup>+</sup>, CD8<sup>+</sup>, CD19<sup>+</sup> and CD56<sup>+</sup> (NK) in patients and control groups.

Table (15) Analysis of difference between surface markers of the intrahepatic lymphocytes (IHL) in patients and control groups (unpaired t-test):

| Surface marker<br>(%)                     | Patients (N=21) Mean ± SD | Control<br>(N=20)<br>Mean ± SD | t value | P value |
|---|---------------------------|--------------------------------|---------|---------|
| - T – cell                                |                           |                                |         |         |
| CD3 <sup>+</sup>                          | 61.23 ± 14.29             | 60.34 ± 12.16                  | 0.218   | 0.828   |
| CD4 <sup>+</sup>                          | 18.07 ± 8.54              | 10.3 ± 3.53                    | 3.923   | 0.001   |
| CD8 <sup>+</sup>                          | 43.19 ± 10.29             | 50.36 ± 10.36                  | -2.223  | 0.032   |
| CD4 <sup>+</sup> / CD8 <sup>+</sup> ratio | 0.43 ± 0.21               | $0.20 \pm 0.06$                | 4.020   | 0.001   |
| -B- cells (CD19 <sup>+</sup> )            | 5.15 ± 2.7                | 3.78 ± 1.79                    | 1.903   | 0.065   |
| - NK cells                                | 24.97 ± 7.38              | 18.27 ± 3.16                   | 3.72    | 0.001   |

There was a statistically significant difference between the proportion of intrahepatic CD4<sup>+</sup>, CD8<sup>+</sup>, CD4<sup>+</sup> / CD8<sup>+</sup> ratio and NK lymphocytes in patients and control groups.

There was no statistically significant difference between the proportion of intrahepatic CD3<sup>+</sup> & CDI9<sup>+</sup> lymphocytes in both groups.

Table (16) Analysis of difference between surface markers of peripheral blood lymphocytes (PBL) and intrahepatic lymphocytes (IHL) in patients group (Paired t-test):

| Patients (N=21)                     |                  |                  |         |         |  |  |
|-------------------------------------|------------------|------------------|---------|---------|--|--|
| Surface marker (%)                  | PBL<br>Mean ± SD | IHL<br>Mean ± SD | t value | P value |  |  |
| - T – cell                          |                  |                  |         |         |  |  |
| CD3 <sup>+</sup>                    | 65.64 ± 9.5      | 61.2 ± 14.2      | 1.688   | 0.107   |  |  |
| CD4 <sup>+</sup>                    | 43.95 ± 7.3      | 18.06 ± 8.5      | 18.416  | 0.001   |  |  |
| CD8 <sup>+</sup>                    | 22.12 ± 8.3      | 43.19 ± 10.3     | 3.663   | 0.001   |  |  |
| CD4 <sup>+</sup> / CD8 <sup>+</sup> | 2.38 ± 1.11      | $0.4 \pm 0.22$   | 8.33    | 0.001   |  |  |
| -B- cells (CD19 <sup>+</sup> ).     | 6.17 ± 4.1       | 5.15 ± 2.7       | 1.12    | 0.12    |  |  |
| - NK (CD56 <sup>+</sup> )           | 14.02 ± 5.03     | 24.97 ± 7.38     | 5.08    | 0.001   |  |  |

There was a significant positive difference between intrahepatic (IHL) and peripheral blood (PBL) CD4+, CD8+, CD4<sup>+</sup> / CD8<sup>+</sup> ratio and NK cells proportions among the patients group (P < 0.05). There was no significant difference between (PB) & (IH) CD3+, CD19+ in patients group (P > 0.05).

Table (17) Correlation between surface markers of the peripheral blood lymphocytes (PBL) and histopathological activity of hepatitis C evaluated by Knodell's score:

| Surface marker of | Knodell's score |         |  |  |
|-------------------|-----------------|---------|--|--|
| (PBL)             | <b>r</b> .      | P       |  |  |
| CD3               | 0.598           | 0.004*  |  |  |
| CD4               | 0.763           | 0.000** |  |  |
| CD8               | -0.002          | 0.993   |  |  |
| CD4/ CD8 ratio    | 0.213           | 0.353   |  |  |
| CD19              | 0.230           | 0.316   |  |  |
| CD56              | -0.250          | 0.275   |  |  |

There was a statistically significant positive correlation between the peripheral blood CD3, CD4 and Knodell's score in patients group (P <0.05). There was no statistically significant correlation between knodell's score and blood CD8, CD19 and CD56 in patients group.

Table (18) Correlation between surface markers of intrahepatic lymphocytes (IHL) and histopathological activity of hepatitis C (HAl) evaluated by Knodell's score:

| Surface marker | Knodel | l's Score |
|----------------|--------|-----------|
| (IHL)          | r      | P         |
| CD3            | 0.577  | 0.006*    |
| CD4            | 0.894  | 0.000**   |
| CD8            | 0.056  | 0.809     |
| CD4/ CD8 ratio | 0.840  | 0.000**   |
| CD19           | -0.040 | 0.864     |
| CD56           | 0.155  | 0.502     |

There was a statistically significant positive correlation between the intrahepatic CD4, CD4 / CD8 ratio and CD3 and the Knodell's score in patients group. There was no statistically significant positive correlation between intrahepatic CD8, CD19 and CD56 and Knodell's score in patients group.

Table (19) Correlation between ALT & AST levels and surface markers of the peripheral blood lymphocytes (PBL) in patients group:

| Surface marker of | ALT (U/L) |        | AST (U/L) |       |
|-------------------|-----------|--------|-----------|-------|
| (PBL)             | R         | р      | r         | P     |
| CD3               | -0.245    | -0.284 | -0.773    | 0.096 |
| CD4               | -0.065    | 0.779  | -0.278    | 0.223 |
| CD8               | -0.212    | 0.357  | -0.129    | 0.576 |
| CD4/ CD8 ratio    | 0.148     | 0.523  | -0.042    | 0.857 |
| CD19              | -0.296    | 0.192  | -0.516    | 0.067 |
| CD56              | -0.126    | 0.587  | -0.046    | 0.843 |

No significant correlation was found between ALT & AST levels and the surface markers of the blood lymphocytes.

Table (20) Ccorrelation between ALT & AST levels and surface markers of intrahepatic lymphocytes in patients group:

| Surface marker of | ALT (U/L) |       | AST    | (U/L) |
|-------------------|-----------|-------|--------|-------|
| (IHL)             | R         | р     | r      | P     |
| CD3               | -0.107    | 0.643 | -0.254 | 0.266 |
| CD4               | 0.018     | 0.938 | -0.233 | 0.309 |
| CD8               | -0.165    | 0.475 | -0.161 | 0.487 |
| CD4/ CD8 ratio    | 0.091     | 0.696 | -0.138 | 0.551 |
| CD19              | -0.367    | 0.102 | -0.493 | 0.063 |
| CD56              | -0.289    | 0.203 | -0.124 | 0.593 |

No significant correlation was found between the levels of ALT & AST and the of intrahepatic lymphocytes in patient group.

Table (21):Descriptive data of the control group:

| Parameter             | Mean ± SD        | Ra   | ange | Unit          |
|-----------------------|------------------|------|------|---------------|
|                       | 1110000          | Min  | Max  |               |
| Age                   | 49.2 ± 8.54      | 37   | 64   | Year          |
| S. Protein            | $7.53 \pm 0.39$  | 6.9  | 8.1  | gm/dl         |
| S. Albumin            | 4.43 ± 0.44      | 3.8  | 5.0  | gm/dl         |
| ALT                   | 33.6 ± 8.56      | 22   | 45.0 | u/l           |
| AST                   | 28.30 ± 6.96     | 20   | 40.0 | u/l           |
| Prothrombin time      | 13.3 ± 0.23      | 12.7 | 14.2 | Seconds       |
| WBcs                  | 6.24 ± 1.76      | 4.1  | 10.0 | Thousands/cmm |
| RBcs                  | 4.98 ± 0.39      | 4.4  | 5.8  | Million/cmm   |
| Hb                    | 13.5 ± 1.50      | 12.1 | 16.1 | gm/dl         |
| Platelets             | 203 ± 22.99      | 175  | 250  | Thousands/cmm |
| Lymphocytes           | $1.6 \pm 0.56$   | 1.0  | 2.7  | Thousands/cmm |
| CD3 (Blood)           | 60.86 ± 4.78     | 51.6 | 70.1 | %             |
| CD4 (Blood)           | 35.7 ± 3.34      | 31.5 | 41.1 | %             |
| CD8 (blood)           | 25.28 ± 3.69     | 19.8 | 30.5 | %             |
| CD4/CD8 ratio (Blood) | $1.45 \pm 0.27$  | 1.7  | 2.1  | %             |
| CD19 (Blood)          | 5.97 ± 3.14      | 2.5  | 9.1  | %             |
| CD56 (Blood)          | 12.64 ± 3.59     | 7.2  | 20.1 | %             |
| CD3 (IH)              | 60.33 ± 12.16    | 40.3 | 77   | %             |
| CD4 (IH)              | 10.13 ± 3.53     | 4.5  | 17.1 | %             |
| CD8 (IH)              | 50.36 ± 10.36    | 35.1 | 63.5 | %             |
| CD4/CD8 ratio (IH)    | $0.200 \pm 0.06$ | 0.1  | 0.3  | %             |
| CD19 (IH)             | $3.78 \pm 1.8$   | 1    | 7.4  | %             |
| CD56 (IH)             | 18.27 ± 3.16     | 12.2 | 26.8 | %             |

Table (22) Descriptive data of the patients group:

| Parameter             | Mean ± SD        | R    | ange | Unit          |
|-----------------------|------------------|------|------|---------------|
|                       | Wioun 2 5g       | Min  | Max  |               |
| Age                   | 47.4 ± 10.67     | 35   | 64   | Year          |
| S. Protein            | $7.77 \pm 0.48$  | 6.3  | 8.5  | gm/dl         |
| S. Albumin            | $4.7 \pm 0.7$    | 2.3  | 5.7  | gm/dl         |
| S. Bilirubin          | 0.91 ± 0.35      | 0.5  | 1.9  | Mg/e          |
| ALT                   | 74.7 ± 17.4      | 40   | 109  | u/e           |
| AST                   | $35.6 \pm 20.5$  | 20   | 48   | u/l           |
| Prothrombin time      | $13.1 \pm 0.98$  | 12   | 15   | Seconds       |
| Prothrombin Conc.     | 83.9 ± 13.3      | 61   | 100  | %             |
| Wbcs                  | $6.5 \pm 2.6$    | 1.7  | 12.3 | Thousands/cmm |
| Rbes                  | $3.9 \pm 0.4$    | 3.2  | 4.7  | Million/cmm   |
| Platelets             | 211.6 ± 69.1     | 85   | 329  | Thousands/cmm |
| Hb                    | 12.2 ± 2.4       | 6.7  | 16   | gm/dl         |
| Lymphocytes           | $1.5 \pm 0.55$   | 0.8  | 3    | Thousands/cmm |
| CD3 (Blood)           | $65.65 \pm 9.53$ | 46.1 | 83   | %             |
| CD4 (Blood)           | 43.96 ± 7.33     | 26.1 | 56.7 | %             |
| CD8 (blood)           | 22.13 ± 8.32     | 10.2 | 36   | %             |
| CD4/CD8 ratio (Blood) | 2.38 ±1.11       | 1    | 4    | %             |
| CD19 (Blood)          | 6.17 ± 3.14      | 4.4  | 10.8 | %             |
| CD56 (Blood)          | $14.02 \pm 5.04$ | 4.6  | 26.3 | %             |
| CD3 (IH)              | 61.24 ± 14.29    | 41   | 86   | %             |
| CD4 (IH)              | 18.06 ± 8.54     | 10.0 | 35.0 | %             |
| CD8 (IH)              | 43.19 ± 10.29    | 28   | 64   | %             |
| CD4/CD8 ratio (IH)    | $0.43 \pm 0.22$  | 0.2  | 0.96 | %             |
| CD19 (IH)             | 5.15 ± 2.73      | 1.5  | 10.5 | %             |
| CD56 (IH)             | 24.97 ± 7.38     | 18   | 33.4 | %             |
| Knodell score         | 4.7 + 3.4        | 1    | 13   | %             |



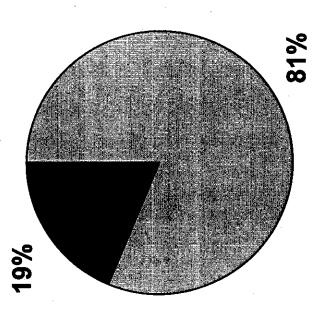
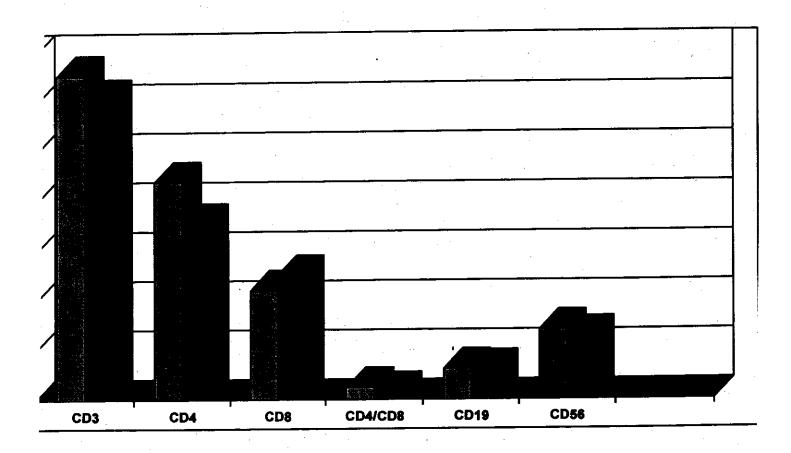


Fig. (25): Sex distribution in patient group.

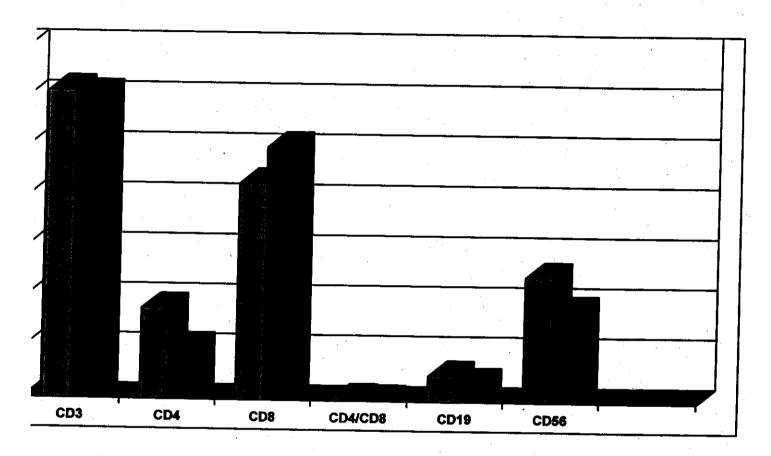




Fig(26) Peripheral blood lymphocytes in patients and control groups.

There was a statistically significant difference between the proportion of PBL CD4<sup>+</sup> and CD4<sup>+</sup>/CD8<sup>+</sup> in patients and control groups (P < 0.05). While, there was no statistically significant difference between the proportion of CD3+, CD8<sup>+</sup>, CD19<sup>+</sup> and CD 56<sup>+</sup> (NK) cells in patients and control groups (P > 0.05).





 $Fig(27)\ Intrahepatic\ lymphocytes\ in\ patients\ and\ control\ groups$ 

There was a statistically significant difference between the proportion of intrahepatic CD4+, CD8+, CD4+/CD8+ ratio and NK lymphocytes in patients and control groups (P < 0.05). There was no statistically significant difference between the proportion of intrahepatic CD3+ and CD 19+ lymphocytes in patients and control group (P. > 0.05).



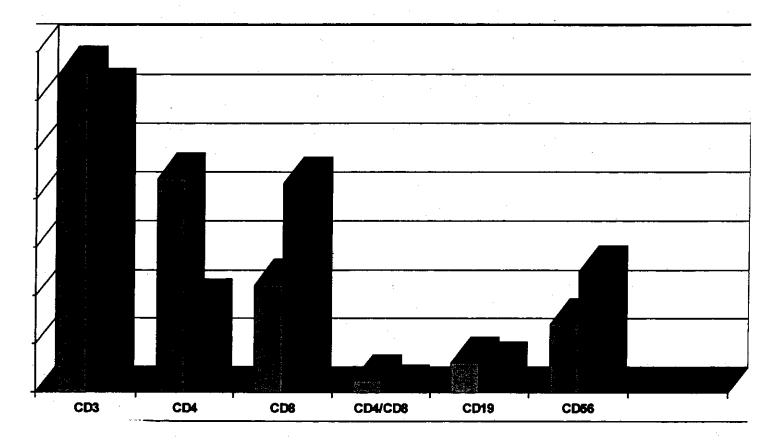


Fig. (28): Peripheral blood and intrahepatic lymphocytes in patients group.

There was a statistically significant difference between the intrahepatic and peripheral blood CD4+, CD8+ and NK cells among patients group (P < 0.05). There was no statistically significant difference between the peripheral blood and intrahepatic CD3+, CDI9+ among patients group (P.> 0.05).

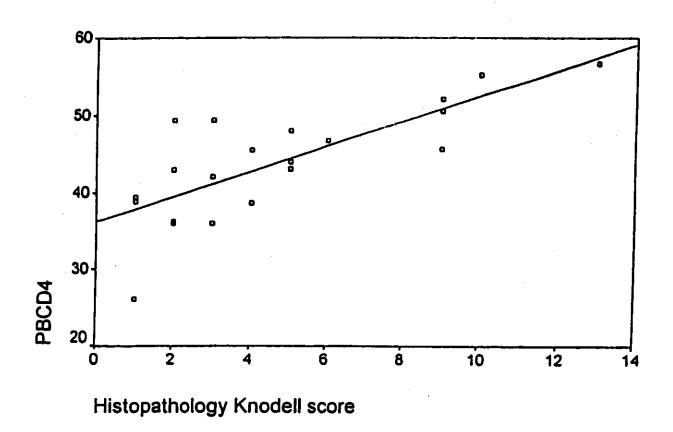


Fig. (29): Correlation between peripheral blood CD4 lymphocytes and Knodell's score in patients group.

There was a statistically significant positive correlation between Knodell's score & CD4+ in patients group (P < 0.05). There was no statistically significant correlation between Knodell's score and CD8+ CD19+, CD56+ (PBL) in patients group.

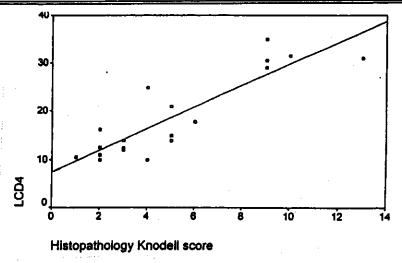


Fig. (30): Correlation between intrahepatic CD4+ lymphocytes and Knodell's score in patients group.

There was a statistically significant positive correlation between Knodell's score & IH CD4, in patients group (P < 0.05). There was no significant correlation between Knodell's score and CD8+ CD19+, CD56+ (IHL) in patients group.

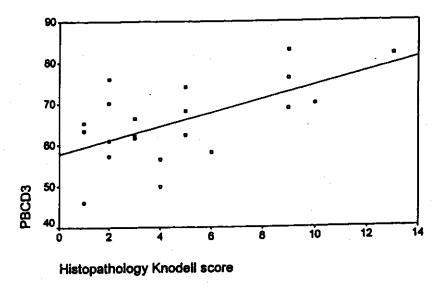


Fig. (31): Correlation between peripheral blood CD3 lymphocytes and knodell's score in patients group.

There was a statistically significant positive correlation between Knodell's score & CD3+ PBL in patients group (P < 0.05).

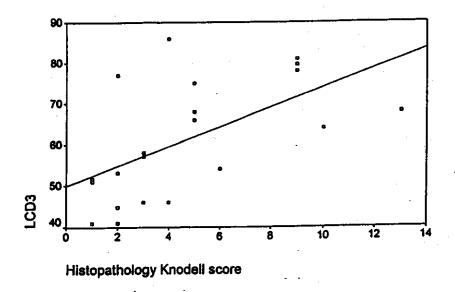


Fig. (32): Correlation between intrahepatic CD3 lymphocytes and Knodell's score in patients group.

There was a statistically significant positive correlation between Knodell's score & CD3+ IHL in patients group (P < 0.05).

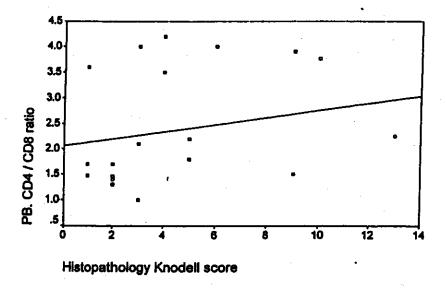


Fig. (33): Correlation between peripheral blood CD4<sup>+</sup>/CD8<sup>+</sup> lymphocytes and Knodell's score in patients group.

No statistically positive correlation was found between peripheral blood CD4<sup>+</sup>/CD8<sup>+</sup> ratio and Knodell's score in patients.

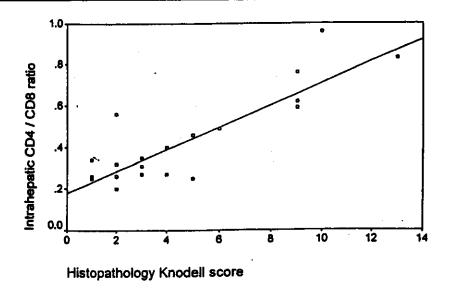


Fig. (34): Correlation between intrahepatic CD4<sup>+</sup>/CD8<sup>+</sup> lymphocytes and Knodell's score in patients group.

There was statistically positive correlation between IH CD4<sup>+</sup>/CD8<sup>+</sup> ratio and Knodell's score in patients.