

The Role Of Adjuvant External Beam Radiotherapy In The Management Of Advanced Prostate Cancer: A Single Institutional Retropective Study In Aba, South Eastern Nigeria.

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Abstract

Prostate cancer is the most common cancer among African American's.

It is rarely diagnosed before the age of fifty years and when diagnosed earlier, it is often due to hereditary factors.

The objective of this study was to review the role of adjuvant external beam radiotherapy in the Management of advanced prostate cancers.

This study was a retrospective study spanning Five years from January 2015 to December 2019.

About 144 Histopathologically diagnosed Adenocarcinoma of the prostate were reviewed within this Period.

Out of these 144 cases, 102 (70.8%) presented for treatment and were commenced on Androgen Deprivation therapy after investigative staging and counseling.

Out of the 102 cases that had Androgen Deprivation therapy (ADT), only 60 persons (58.8%) had follow up External beam radiotherapy (RT) while 42 (41.2%) did not have follow up RT.

Finding showed that those who had ADT alone without follow up RT had shorter time interval before onset of biochemical failure and Castration resistance compared to those who had follow up external Beam radiotherapy.

Findings also showed that the more aggressive the tumor with respect to the Gleason grading, the Shorter the interval before the onset of biochemical failure and castration resistance.

External beam radiotherapy was found to be a very useful adjunct treatment in the management of advanced prostate cancer in Abia increasing the symptom free interval before the onset of castration Resistance.

Keywords: Advanced prostate cancer, external beam radiotherapy, biochemical failure, castration resistance and Aba, Nigeria

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I. Introduction

Prostate cancer is the most common cancer among African American men.

Men of African ancestry have higher incidence, morbidity and mortality.

Due to the poor screening knowledge and practice, most of these cancers present at advanced stage (Mohammed and Mohammed 2011)

Advanced prostate cancer includes any cancer with extra prostatic spread such as:

- Locally advanced disease
- Metastatic prostate cancer
- Castration resistant prostate cancer

Whereas curative radiotherapy is a recommended modality of treatment for organ confined prostate cancers especially when it is high grade.

Similarly, Brachytherapy is a recommended treatment protocol for organ confined disease.

Palliative external beam radiotherapy is useful in advanced prostate cancer with extra prostatic spread.

It is used as

- Part of initial adjuvant treatment of advanced disease 6 months after ADT.
- To improve the symptoms of the disease such as Hematuria, bone pains and obstructive voiding.
- To improve symptom free survival.

In locally advanced and metastatic settings, patients should ideally be managed by Androgen Deprivation therapy for about 6 months before palliative external beam radiotherapy to the prostate.

This interval of 6 months enables metastatic cells to wither or undergo Apoptosis before radiating the prostate. After radiation Therapy, Androgen Deprivation Therapy is continued for another 2 yrs to prevent escaped cells from recovering.

In the castration resistance setting with metastases, extra prostatic cells are better managed with Docetaxel – prednisolone therapy so as to control and localize the disease before application of radiotherapy.

But in a castration resistant setting with localized disease Abiraterone therapy may be preferable before administration of radiotherapy.

There is a place for hypo fractionated radiotherapy giving the opportunity for fewer but larger doses of radiotherapy compared to conventional radiotherapy.

The shorter treatment time improves cost effectiveness, reduces time of exposure and thereby reducing adverse effects and at the same time maintaining good outcome.

II. Methodology

This was a retrospective review of the histo pathologically diagnosed prostate cancers within the study period of 5 yrs between January 2015 to December 2019.

Their case files were withdrawn and relevant information collated from them.

Such relevant information included their personal Data, Histo pathology reports with Gleason grades, Type of ADT giving time of administration of radiotherapy if any and the date of onset Biochemical failure and castration resistance.

These information and data were collated, analyzed and interpreted.

Inclusion Criteria

Men who were diagnosed of prostate cancer and received treatment within the study period were part of the study.

Exclusion Criteria

Men who were diagnosed of prostate cancer within the study period but did not receive any form of treatment were excluded from the study.

III. Results

Table 1- Showing The Age Group Distribution Of The Histopathologically Confirmed Prostate Cancers (ADENOCARCINOMA) N=144

| S/NO | Age Group In Years | Number | Percentage |
|------|--------------------|--------|------------|
| 1 | 40-50yrs | 2 | 1.4% |
| 2 | 51-60yrs | 14 | 9.7% |
| 3 | 61-70yrs | 49 | 34.0% |
| 4 | 71-80yrs | 59 | 40.9% |
| 5 | 81-90yrs | 19 | 13.2% |
| 6 | 91-110yrs | 1 | 0.7% |
| 7 | Total | 144 | 100% |

Fig. 1 Bar Chart

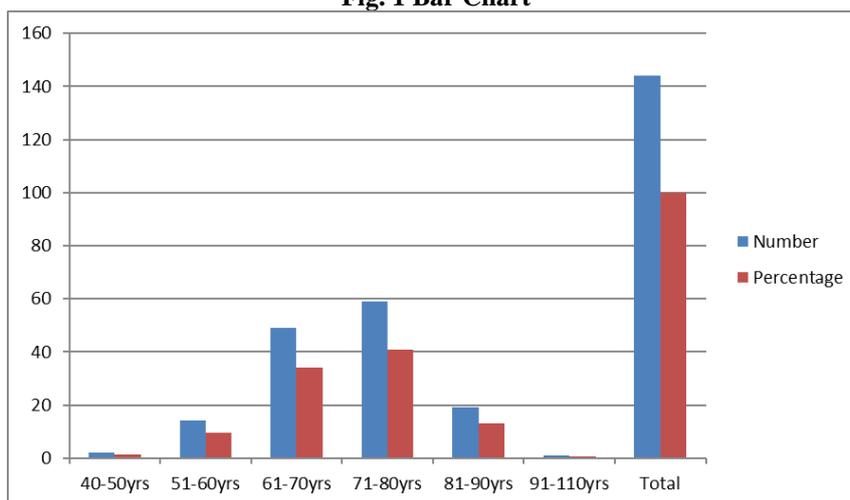


Table 2- Showing The Gleason Grading Groups Of The Histopathologically Confirmed Prostate Cancers N-144

| S/NO | Gleason Score And Sun | Grade | Number | Percentage |
|------|-----------------------|--------------|--------|------------|
| 1 | 3+3=6/10 | Low | 11 | 7.6% |
| 2 | 3+4=7/10 | Intermediate | 21 | 16.7% |
| 3 | 4+3=7/10 | High | 19 | 13.2% |
| 4 | 4+4=8/10 | High | 35 | 24.3% |
| 5 | 4+5=9/10 | High | 18 | 12.5% |
| 6 | 5+4=9/10 | High | 24 | 16.6% |
| 7 | 5+5=10/10 | High | 16 | 11.1% |
| 8 | Total | | 144 | 100% |

Fig. 2 Bar Chart

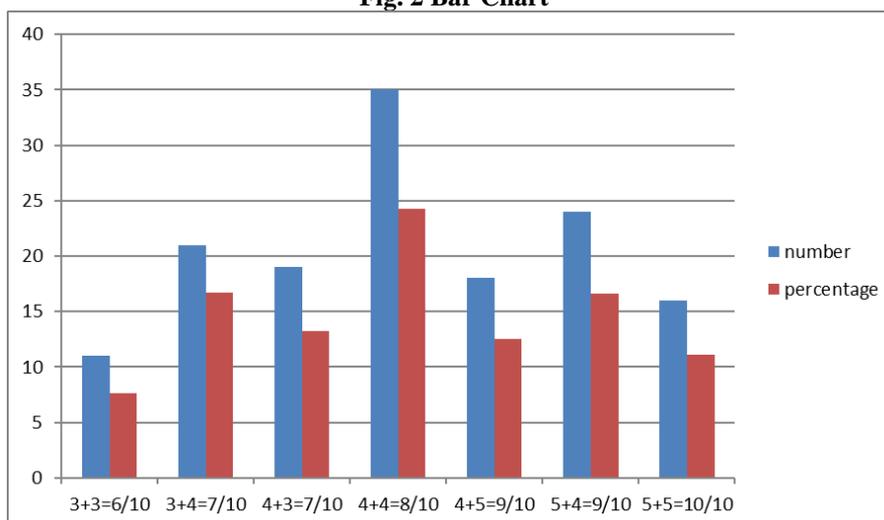


Table 3- Showing The Tumor Grades And Those Who Received Androgen Deprivation Therapy Adt N=102

| S/NO | Tumor Grade | ADT | Number | Percentage |
|------|------------------|-----|--------|------------|
| 1 | 2+3 Low Grade | ADT | 8 | 7.8% |
| 2 | 3+4 Intermediate | ADT | 11 | 10.8% |
| 3 | 4+3 High Grade | ADT | 13 | 12.7% |
| 4 | 4+4 High Grade | ADT | 28 | 27.5% |
| 5 | 4+5 High Grade | ADT | 12 | 11.8% |
| 6 | 5+4 High Grade | ADT | 16 | 15.8% |
| 7 | 5+5 High Grade | ADT | 14 | 13.7% |
| 8 | Total | | 102 | 100% |

Fig. 3 Bar Chart

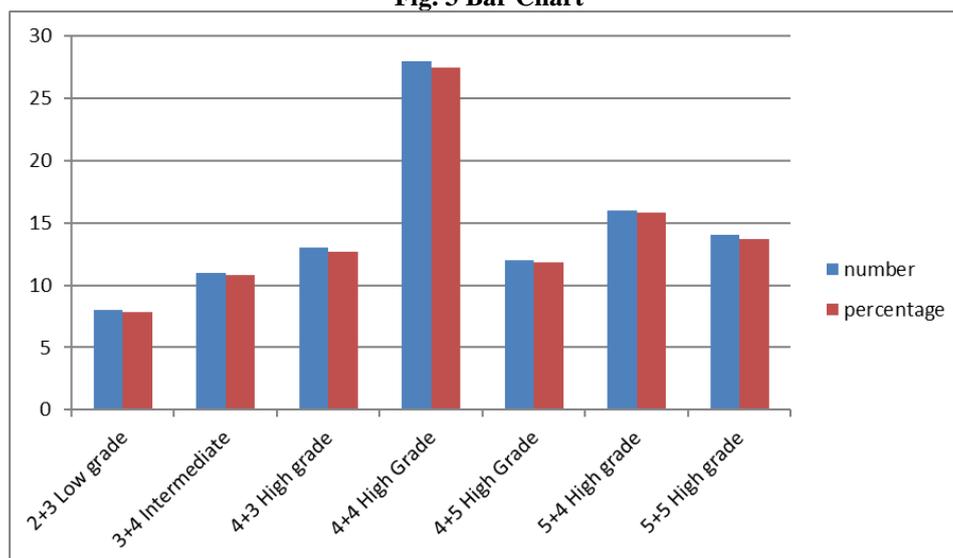


Table 4 Showing The Tumor Grade And Those Who Had Androgen Deprivation Therapy (Adt) With Follow Up Adjuvant External Beam Radiotherapy (Rt)

| S/NO | Gleason Grade | Adt | Number | External Beam Rt | Number |
|------|------------------|-----|--------|------------------|--------|
| 1 | 3+3 Low Grade | Adt | 8 | Rt Radiotherapy | 3 |
| 2 | 3+4 Intermediate | Adt | 11 | Rt Radiotherapy | 6 |
| 3 | 4+3 High Grade | Adt | 13 | Radiotherapy | 7 |
| 4 | 4+4 High Grade | Adt | 28 | Radiotherapy | 18 |
| 5 | 4+5 High Grade | Adt | 12 | Radiotherapy | 7 |
| 6 | 5+4 High Grade | Adt | 16 | Radiotherapy | 9 |
| 7 | 5+5 High Grade | Adt | 14 | Radiotherapy | 10 |
| 8 | Total | | 102 | | 60 |

Table 5 Showing Cases That Had Androgen Had Androgen Deprivation Therapy (Adt) Without Follow Up External Beam Radiotherapy (Rt)

| S/NO | Tumor Grade | Adt | Number | Nil External Beam Radiotherapy | Number |
|------|-----------------------|-----|--------|--------------------------------|--------|
| 1 | 3+3=10 High Grade | Adt | 8 | No Follow Up Radiotherapy | 5 |
| 2 | 3+4=7/10 Intermediate | Adt | 11 | No Follow Up Radiotherapy | 5 |
| 3 | 4+3=7/10 High Grade | Adt | 13 | No Follow Up Radiotherapy | 6 |
| 4 | 4+4=8/10 High Grade | Adt | 28 | No Follow Up Radiotherapy | 10 |
| 5 | 4+5 High Grade | Adt | 12 | Nofollow Up Radiotherapy | 5 |
| 6 | 5+5 High Grade | Adt | 16 | No Follow Up Radiotherapy | 7 |
| 7 | 5+5 | Adt | 14 | No Follow Up Radiotherapy | 4 |
| | Total | | 102 | No Follow Up Radiotherapy | 42 |

Table 6 Showing Cases Who Received Adt Only And Those With Both Adt Follow Up Radiotherapy

| S/No | Gleason Grade | Therapy Recieved | Number | Total |
|------|----------------------------------|------------------|--------|-------|
| 1 | 3+3=6/10 Low Grade | Adt Only | 5 | 8 |
| 2 | 3+4=9/10 High Grade Intermediate | Adt Only | 6 | 11 |
| | | Adt And Rt | 5 | |
| 3 | 4+3=7/10 High Grade | Adt Only | 7 | 13 |
| | | Adt And Rt | 6 | |
| 4 | 4+4 High Grade | Adt Only | 18 | 28 |
| | | Adt And Rt | 10 | |
| 5 | 4+5 High Grade | Adt Alone | 7 | 12 |
| | | Adt And Rt | 5 | |
| 6 | 5+5 High Grade | Adt Alone | 10 | 14 |
| | | Adt And Rt | 4 | |

Table 7 Showing Average Time Before Onset Of Biochemical Failure And Castration Resistance In Years

| S/No | Gleason Grade | Therapy Employed | Average Time Before Resistance |
|------|-----------------------|-------------------|--------------------------------|
| 1 | 3+3=6/10 Low Grade | Adt Alone Adt+Rt | <3yrs Nil |
| 2 | 3+4=7/10 Intermediate | Adt Alone Adt+Rt | <3yrs Nil |
| 3 | 4+3=7/10 High Grade | Adt Alone Adt +Rt | <2yrs >8yrs |
| 4 | 4+4=8/10 High Garde | Adt Alone Adt+Rt | <1 ½ Yrs >8yrs |
| 5 | 4+5=9/10 High Grade | Adt Alone Adt+Rt | <1 ½ Yrs >7yrs |
| 6 | 5+4=9/10 | Adt Alone Adt+Rt | <1 ½ Yrs >8 Yrs |
| 7 | 5+5=10/10 | Adt Alone Adt+Rt | <1 Yrs >6 Yrs |

IV. Discussion

Advanced prostate cancer refers to cancer with extra prostatic extension or one that has undergone castration resistance.

Even though advanced, palliative radiotherapy is an effective tool for management.

Palliative external radiotherapy is used for

- Managing symptoms in advanced prostate cancer particularly bone pains, haematuria, urinary obstruction and rectal complications.
- Extending symptom free survival
- Prolonging the interval before onset of biochemical failure and castration resistance.

In our study, we found the above uses in men who had follow up adjuvant external beam radiotherapy following primary treatment with androgen deprivation therapy.

These patients experienced

- Long symptom free interval
- Delayed onset of biochemical failure and castration resistance
- Extended survival.

Francis Asamoah et al, in Ghana, in their work on contemporary radiation treatment on prostate cancer in Africa- A Ghanaian experience, found the biochemical disease free survival rates (bDFS) for low, intermediate and high risk groups were 95%, 70% and 48% respectively.

Both Gleasn score and prostate specific antigen (PSA) were significant predictors for biochemical disease free survival rates.

They concluded that majority of the prostate cancers were locally advanced cancers at the time of presentation for radiotherapy.

bDFS was significantly better for low and intermediate risk diseases than high risk disease.

Rose J. N et al, in their work on the role of radiation therapy in the treatment of the metastatic castration resistant prostate cancer, found out that

- Localized castration resistant prostate cancer can be associated with significant morbidity.
- High dose palliative radiotherapy can be a useful palliative maneuver to obtain local control
- High dose palliative radiotherapy to the prostate is associated with minimal significant toxicity.
- Extended survival is possible in this prostate cancer sub group.

They concluded that high dose palliative radiotherapy was well tolerated and provided high rate of local control in clinically localized castration resistant prostate cancer population.

Although prostate cancer remained the most frequent cause of death, some patients had extended survival without evidence of disease progression.

V. Conclusion

Adjuvant external beam radiotherapy after pre-treatment with androgen deprivation therapy is an important adjunct treatment in Aba.

It was found to delay onset of biochemical failure and castration resistance and generally had extended symptom free survival.

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